Database ID Gene Symbol		PubMED ID	Annotation
131550	Id2	25902484	Group 3 innate lymphoid cells (ILC3s) can mediate immune surveillance, which constantly maintains a proper microbiota, to facilitate early colonization resistance through an Id2-dependent regulation of Il22. Iltifb (Il22b) increases the TNF-alpha-dependent induction and secretion of several immune-modulatory molecules such as initial complement factors, antimicrobial peptides and chemokines in primary
192083	Iltifb	21469124	keratinocytes. Iltifb-mediated induction of innate immunity is crucial for the maintenance of epidermal integrity during infection with Candida albicans. (Demonstrated in human) Iltifb (Il22) is produced by lymphoid tissue-inducer cells where it regulates the maintenance of colonic lymphoid structures during Citrobacter rodentium infection, a mechanism that bridges the lymphotoxin
192083	Iltifb	21874025	pathway to mucosal epithelial defense mechanisms. ILTIFB (IL22) protects intestinal stem cells from
192083	Iltifb	22921121	inflammatory tissue damage and regulates sensitivity to graft versus host disease. (Demonstrated in human) Flagellin induces Tlr5-dependent Il22 production and Nlrc4-dependent Il18 production to promote a protective gene expression program in intestinal
192083	Iltifb	25395539	epithelial cells and elimination of rotavirus-infected cells. Cxcl16-Cxcr6 crosstalk coordinates the intestinal
192083	Iltifb	25456160	topography of II22 secretion required for mucosal defence against Citrobacter rodentium infection. II22 augments the expression of II18 mRNA and inactive precursor protein (proIL-18) in intestinal epithelial cells after Toxoplasma gondii or Citrobacter
192083	Iltifb	25680273	rodentium infection and maintains the homeostatic amount of proIL-18 in the ileum. Innate lymphoid cells, potent producers of Il22 after
192083	Iltifb	26649819	intestinal injury, increase the growth of mouse small intestine organoids in an Il22-dependent fashion. Group 3 innate lymphoid cells (ILC3s) can mediate immune surveillance, which constantly maintains a
192083	Iltifb	25902484	proper microbiota, to facilitate early colonization resistance through an Id2-dependent regulation of Il22. The G3BP1-CAPRIN1-PRKRA complex represents a new mode of PRKRA activation and links stress responses with innate immune activation through PRKRA without a requirement for foreign double-
76030	PRKRA	25784705	stranded RNA pattern recognition. A defective interfering RNA isolated from the Hu-191
76030	PRKRA	26608320	vaccine strain of measles virus is sensed by PRKRA and DDX58 to initiate an innate antiviral response.

## Supplementary Table 4. Detailed information of the immune-related genes in InnateDB database.

76030	PRKRA	26454002	DDX3X participates in antiviral innate immunity by controlling translation of PRKRA.
229341	DDX3X	18583960	DDX3X is a critical component of the TANK-binding kinase 1 (TBK1)-dependent innate immune response. DDX3X is an antiviral MAVS (IPS-1) enhancer, where
229341	DDX3X	20127681	it can bind viral RNA to join it in the MAVS complex and this augments virus-mediated IFN-beta induction and host cell protection against virus infection. DDX3X RNA helicase can augment TBK1/IKKepsilon activity and hepatitis B virus (HBV) polymerase (HBV Pol) inhibits this TBK1/IKBKE activity by disrupting
229341	DDX3X	20657822	the interaction between IKBKE and DDX3X, providing an explanation for how HBV evades innate immune response in the early phase of the infection. DDX3X initiates a multifaceted cellular program involving dynamic associations with hepatitis C virus (HCV) RNA and proteins, CHUK, stress granules, and
229341	DDX3X	25740981	lipid droplet surfaces for its crucial role in the HCV life cycle.
229341	DDX3X	26454002	DDX3X participates in antiviral innate immunity by controlling translation of PRKRA. DDX58 (RIG-I) is a cytoplasmic RNA helicase that
55854	DDX58	15208624	functions as an intracellular sensor of dsRNA leading to the induction of Interferon (IFN) production independently of TLR signalling. DDX58 (RIG-I) first and second caspase recruitment domains (CARDs) have distinct roles in TRIM25-
55854	DDX58	18948594	mediated RIG-I ubiquitination, which leads to initiation of an antiviral signalling cascade. DDX58 serves as a critical link between TLR3 and
55854	DDX58	19074283	type-II-IFN signalling pathways in innate antiviral immune responses. DDX58 plays an essential role in Toll-like receptor
55854	DDX58	19683681	<ul><li>(TLR)-stimulated phagocytosis, demonstrating that DDX58 plays a role not only in antiviral responses but in antibacterial responses as well.</li><li>DDX58 plays a key role in the expression of TNF- alpha in macrophages in response to LPS stimulation,</li></ul>
55854	DDX58	18523264	mainly for the late phase LPS-induced expression of TNF-alpha. DDX58 is a sensor able to activate the inflammasome
55854	DDX58	19915568	<ul> <li>in response to certain RNA viruses by binding to the adaptor PYCARD to trigger the caspase-1 (CASP1)-dependent inflammasome activation and IL-1-beta production.</li> <li>DDX58 binds specifically to K63-polyubiquitin chains through its tandem caspase recruitment domains (CARDs) that act as a ubiquitin sensor in a manner that depends on RNA and ATP, demonstrate that un-</li> </ul>
55854	DDX58	20403326	anchored K63-polyubiquitin chains are signalling molecules in antiviral innate immunity.

			DDX58 (RIG-I) phosphorylation on serine 8 operates as a negative switch of RIG-I activation by suppressing
55854	DDX58	20406818	TRIM25 interaction. DDX58 innate immune response to viral infection of
			human cells is modified by a functional polymorphism
55854	DDX58	20511549	in the RIG-I caspase recruitment domain (CARD). DDX58 (RIG-I) is responsible for the cytosolic recognition of Legionella pneumophila RNA and the subsequent induction of type I IFN response.
55854	DDX58	19936053	(Demonstrated in murine model) DDX58 and NOD2 colocalize to cellular ruffles and cell-cell junctions to form a protein complex via the CARD domains. DDX58 negatively regulates ligand- induced NFkB signalling mediated by NOD2, and conversely, NOD2 negatively regulates type I
55854	DDX58	21690088	interferon induction by DDX58. DDX58, through the TRAIL pathway, initiates apoptosis in hepatocytes infected with hepatitis C Virus to suppress viral replication. HCV envelope proteins counteract the antiviral host defence by
55854	DDX58	21695051	inhibiting the expression of DDX58. DDX58 (RIG-I) ubiquitination is inhibited by arterivirus and nairovirus deubiquitinating enzymes (DUBs), resulting in the inhibition of RIG-I-like receptor (RLR)-mediated innate immune signalling.
55854	DDX58	22072774	(Demonstrated in mice) Antiviral stress granules containing DDX58 (RIG-I) and EIF2AK2 (PKR) have a critical role in viral detection and innate immunity. (Demonstrated in
55854	DDX58	22912779	mouse) DDX58 (RIG-I) stimulation with a synthetic ligand
55854	DDX58	23744645	inhibits HIV replication in macrophages. RNF135 is essential for the association of DDX58
55854	DDX58	23950712	<ul><li>(RIG-I) and TRIM25, resulting in the activation of RIG-I signalling.</li><li>ISG15 does not directly alter human rhinovirus replication but modulates immune signalling via the viral sensor protein DDX58 to impact production of</li></ul>
55854	DDX58	24448099	CXCL10, which has been linked to innate immunity to viruses. Human rhinovirus infection of epithelial cells induces the expression and secretion of ISG15, which modulates immune responses via effects on DDV58
55854	DDX58	24448099	modulates immune responses via effects on DDX58, and by regulating CXCL10 production. The antisense L region of encephalomyocarditis virus
55854	DDX58	24550253	associates with DDX58 and is a key determinant of IFIH1 stimulation of infected cells. IFI16 transcriptionally regulates type I interferons and DDX58 (RIG-I) and controls the interferon response to
55854	DDX58	25002588	both DNA and RNA viruses.

			Paramyxoviruses trigger the DNA-damage response, a pathway required for RPS6KA5 activation of phospho Ser 276 RELA formation to trigger the IRF7-DDX58
55854	DDX58	25520509	amplification loop necessary for mucosal interferon production. DDX58 dually functions as an hepatitis B virus sensor activating innate signalling and as a direct antiviral
55854	DDX58	25557055	factor by counteracting the viral polymerase in hepatocytes. DDX58 is the primary pattern recognition receptor (PRR) for influenza A virus (IAV), but IFIH1 is a significant contributor to the cellular defense against
55854	DDX58	26074083	IAV. Signalling through both DDX58 and TLR3 is important for interferon induction by influenza A virus
55854	DDX58	25880109	in alveolar epithelial cells. Hepatitis B virus-induced MIR146A attenuates cell-
55854	DDX58	27210312	<ul><li>intrinsic anti-viral innate immunity through targeting DDX58 and IFIT3.</li><li>MIR485 exhibits bispecificity, targeting DDX58 in cells with a low abundance of H5N1 virus and viral PB1 in cells with increased amounts of the H5N1 virus.</li></ul>
55854	DDX58	26645583	
			A defective interfering RNA isolated from the Hu-191 vaccine strain of measles virus is sensed by PRKRA
55854	DDX58	26608320	and DDX58 to initiate an innate antiviral response. ATP binding is required for DDX58 signalling on viral RNA. ATP hydrolysis provides an important function
55854	DDX58	26371557	by recycling DDX58 and promoting its dissociation from non-pathogenic RNA. Influenza B virus induces IRF3 activation and IL29 (IFNL1) gene expression without a requirement for
			viral protein synthesis or replication and DDX58 is the critical pattern recognition receptor needed for IRF3
55854	DDX58	26378160	activation. EFTUD2 is a novel innate immune antiviral regulator that restricts hepatitis C virus infection through a DDX58/IFIH1-mediated, JAK-STAT-independent
55854	DDX58	25878102	pathway. MIR136 exhibits potent antiviral activity against H5N1
55854	DDX58	26450567	influenza A virus and acts as an immune agonist of DDX58. MIR136 exhibits potent antiviral activity against H5N1 influenza A virus and acts as an immune agonist of DDX58.
126353	MIR136	26450567	
223927	Mir146	22545247	Mir146 directly targets Relb to modulate the amplitude of monocyte responses to inflammatory challenges.

			Mir146 is a mechanosensitive miRNA that modulates mechanotransduction and pressure-induced inflammation in small airway epithelium.
223927	Mir146	22593544	(Demonstrated in human) Nod2 driven inflammation is regulated by nitric oxide responsive Mir146 that facilitates activation of sonic
223927	Mir146	24092752	hedgehog (SHH) signalling by targeting Numb expression. T-cell-intrinsic Mir155 is required for type-2 immunity, in part through regulation of S1pr1, whereas
223927	Mir146	25024218	T-cell-intrinsic Mir146 is required to prevent overt Th1/Th17 skewing. Mir146 attenuates sepsis-induced cardiac dysfunction by preventing NF Î °B activation, inflammatory cell infiltration, and inflammatory cytokine production via
223927	Mir146	26048146	targeting of Irak1 and Traf6 in both cardiomyocytes and inflammatory monocytic cells. Mir146 has a role in constraining intestinal barrier function, a process that alters gut homeostasis and
223927	Mir146	26456940	increases susceptibility to dextran sodium sulphate- induced colitis. Nod2-deficient mice have impaired resistance to Mycobacterium tuberculosis infection through defective innate and adaptive immunity.
179050	Nod2	18981137	Nod2 functions in non-hematopoietic cells of the small intestinal crypts and this is critical for protecting mice from a Th1-driven granulomatous inflammation in the ilum.
179050	Nod2	20679225	Nod2 and Nod1 account for neutrophil recruitment to the lungs of mice infected with Legionella pneumophila.
179050	Nod2	20685341	Nod2 and Nod1 activation results in substantial secretion of Ccl5 by murine macrophages and induces binding of NF-kappaB subunits to Ccl5 promoter.
179050	Nod2	17705131	Nod2 and Nod1 can detect Legionella pneumophila and
179050	Nod2	21072876	these receptors modulate the in vivo pulmonary immune response differently. Nod2 is both a positive and negative regulator of Tlr4 - the effect it exerts is dependent on the presence of MDP. Nod2 upon engagement with its ligand, MDP, positively regulates Tlr4-mediated signaling; in the absence of MDP, Nod2 negatively regulates the Tlr4 pathway.
179050	Nod2	21199260	patiiway.

170050	NJ 10	2122/705	Nod2 is a peripheral peptidoglycan intracellular sensor and is important for the progression and pathogenesis of experimental autoimmune encephalomyelitis (animal model of multiple sclerosis).
179050	Nod2	21236705	Nod2 detects heat-killed Legionella pneumophila and stimulates NFkB and IFN-beta promoter activity. Nod2 deficiency results in increased proinflammatory cytokine expression at 4hrs and greater neutrophil recruitment to the lung.
179050	Nod2	21108472	Ddx58 and Nod2 colocalize to cellular ruffles and cell- cell junctions to form a protein complex via the CARD domains. Ddx58 negatively regulates ligand-induced NFkB signalling mediated by Nod2, and conversely, Nod2 negatively regulates type I interferon induction
179050	Nod2	21690088	by Ddx58. (Demonstrated in human) Nod2 recognition of muramyl dipeptide, a component of bacterial cell walls, improves the barrier function of
179050	Nod2	22750073	intestinal epithelial cells. Nod2 enhances the innate immune response of alveolar
179050	Nod2	22531915	<ul> <li>macrophages to Mycobacterium tuberculosis in human.</li> <li>(Demonstrated in human)</li> <li>Nod1 and Nod2 synergize with Tlr4 in dendritic cells to increase IL12 production and enhance invariant natural killer T (iNKT) cell activation, and are important regulators of the IFN gamma response by iNKT cells during S. typhimurium and L.</li> </ul>
179050	Nod2	24163408	monocytogenes infections. Salmonella enterica serovar Typhimurium Î mmsbB that possesses a modified lipid A triggers exacerbated colitis in the absence of Nod1 and/or Nod2, which is
179050	Nod2	25423082	likely due to increased Tlr2 stimulation. Nod2 regulates type-1 cytokine responses to Mycobacterium avium but is not required for the
179050	Nod2	25817335	control of M. avium infection in vivo. Leukotriene B4 acts on Nod2 pathway to enhance the
179050	Nod2	26444420	immune response against influenza A infection. Under acidic conditions both pro-inflammatory forms
205910	Illa	24022484	of Il1a and Il1b are regulated independently of the NLRP3 inflammasome. In response to adenovirus infection, the IL1A-IL1R1-CXCR2 signalling axis cooperates with complement to recruit Ly-6G+7/4+ polymorphonuclear leukocytes to the splenic marginal zone (MZ) in the proximity of virus-containing MARCO+ residential MZ
205910	Illa	24651866	macrophages, which are subsequently eliminated. NLRP3 inflammasome formation is dispensable for alum-induced innate immunity but Il1a and Il1b are both necessary for alum-induced neutrophil influx in
205910	Illa	26536497	vivo.

205910	Illa	26439902	Il1a directly senses DNA damage and acts as signal for genotoxic stress without loss of cell integrity. Interleukin-1 (IL1A/IL1B) plays a key role in the interaction between local vessel wall cells and invading
66482	IL1A	25463072	monocytes to multiply cholesterol-triggered inflammation in the vessel wall. IFNG interferes with the IL-1/NFKBIZ axis in $\hat{I}^2$ -glucan-activated dendritic cells and promotes T cell-
66482	IL1A	25474109	mediated immune responses with increased release of IFNG and IL22, and diminished production of IL17A. CASP4 is a critical regulator of noncanonical inflammasome activation that initiates defence against
66482	IL1A	25964352	bacterial pathogens in primary macrophages by mediating cell death and IL1A release IL1A directly senses DNA damage and acts as signal
66482	IL1A	26439902	for genotoxic stress without loss of cell integrity. IFIH1 (MDA5) recognizes distinct RNA viruses and plays a major role in the elimination of RNA viruses in
73750	IFIH1	16785313	vivo.
73750	IFIH1	15563593	IFIH1 binds V proteins of paramyxoviruses and this inhibit its activation of the IFNB1 (IFN-beta) promoter. IFIH1 is a double-stranded RNA-dependent ATPase that contains both a caspase recruitment domain and
			RNA helicase motifs. IFIH1 may also function as a mediator of interferon (IFN)-induced growth inhibition
73750	IFIH1	11805321	and/or apoptosis. IFIH1 is indispensable for sustained expression of IFN in response to paramyxovirus infection and provide the
73750	IFIH1	20107606	first evidence of MDA5-dependent containment of in vivo infections caused by (-) sense RNA viruses. IFIH1 is an RNA helicase and is a key component in activating the expression of type I IFN in response to viral infection. Viral mRNA with 5' cap and 3' poly(A) from parainfluenza virus 5 is able to activate IFN expression through RNASEL-IFIH1 signalling pathway.
73750	IFIH1	21245317	
			IFIH1 (MDA5) is responsible for the cytosolic recognition of Legionella pneumophila RNA and the subsequent induction of type I IFN response.
73750	IFIH1	19936053	(Demonstrated in murine model) (Demonstrated in murine model) IFIH1 deficiency results in a delayed type I IFN and attenuated type III IFN response to rhinovirus infection, leading to a transient increase in viral titer. Upon recognition of viral dsRNA, IFIH1 synergizes with TLR3 to induce pro-inflammatory signals leading to airways inflammation and hyper-responsiveness.
73750	IFIH1	21637773	(Demonstrated in murine model) Paramyxovirus V proteins bind to IFIH1 (MDA5) to
73750	IFIH1	23328395	disrupt viral RNA recognition and induction of antiviral immunity.

			The antisense L region of encephalomyocarditis virus associates with DDX58 and is a key determinant of
73750	IFIH1	24550253	IFIH1 stimulation of infected cells. IFIH1 recognizes hepatitis C virus (HCV) to initiate
73750	IFIH1	25463548	host interferon response during HCV infection. DDX58 is the primary pattern recognition receptor (PRR) for influenza A virus (IAV), but IFIH1 is a significant contributor to the cellular defense against
73750	IFIH1	26074083	IAV. EFTUD2 is a novel innate immune antiviral regulator that restricts hepatitis C virus infection through a DDX58/IFIH1-mediated, JAK-STAT-independent
73750	IFIH1	25878102	pathway. EFTUD2 is a novel innate immune regulator that restricts hepatitis C virus infection through an RIG-
54412	EFTUD2	25878102	I/MDA5-mediated, JAK-STAT-independent pathway. EFTUD2 is a novel innate immune antiviral regulator that restricts hepatitis C virus infection through a DDX58/IFIH1-mediated, JAK-STAT-independent
54412	EFTUD2	25878102	pathway. Tlr2 expression in astrocytes is induced by TNF alpha
154818	Tlr2	18768838	and NF Kappa B-dependent pathways. Tlr2 recognition of Staphylococcal peptidoglycan leads
154818	Tlr2	18621910	to induction of beta-defensin-2. Tlr2 is involved in Respiratory syncytial virus (RSV) recognition and subsequent innate immune activation
154818	Tlr2	19019963	by promoting neutrophil migration and dendritic cell activation within the lung. Tlr4, Tlr2, or Tlr3 cooperate with proteinase-activated receptors (PARs) for activation of nuclear factor-
154818	Tlr2	19865078	kappaB-dependent signaling in mucosal epithelial cell lines. Tlr2 is a signaling receptor that is activated by lipopolysaccharide (LPS) in a response that depends on
154818	Tlr2	9751057	LPS-binding protein and is enhanced by CD14. Tlr2 is a signal transducer for soluble Peptidoglycan
154818	Tlr2	10364168	and lipoteichoic acid (LTA) in addition to LPS. Tlr2 recognizes Gram-positive bacterial cell wall components, leading to the activation of the innate
154818	Tlr2	10384090	immune system. Tlr2 is a molecular link between microbial products,
154818	Tlr2	10426996	apoptosis, and host defense mechanisms. TLR2/MyD88/PI3K/Rac1/Akt pathway mediates the activation of LTA-induced mitogen-activated protein kinases (MAPKs), which in turn initiates the activation of NF-kappaB, and ultimately induces cPLA2/COX-2-
154818	Tlr2	20167866	dependent PGE2 and IL-6 generation. Tlr2 and Tlr4 activate murine macrophages and this is negatively regulated by a Lyn/PI3K module and
154818	Tlr2	20385881	promoted by SHIP1.

			Tlr2 is a molecular link between increased dietary lipid intake and the regulation of glucose homeostasis, via regulation of energy substrate utilization and tissue
154818	Tlr2	20407745	inflammation. Tlr2 activation induces type I interferon responses from endolysosomal compartments where it is translocated
154818	Tlr2	20422028	to upon ligand engagement. Tlr2 and Myd88-dependent phosphatidylinositol 3- kinase and Rac1 activation facilitates the phagocytosis
154818	Tlr2	20368346	of Listeria monocytogenes by murine macrophages. Tlr1 :: Tlr2 dimeric pairs recognize malarial glycosylphosphatidylinositols (GPI) to initiates
154818	Tlr2	21439957	intracellular signalling and the production of pro- inflammatory cytokines. Tlr2 recognizes Thermus aquaticus extracellular polysacchride, YT-1, and induces the production of
154818	Tlr2	21454596	cytokines TNF and IL6 in peritoneal macrophages. Tlr2::Tlr6 synergistically interacts with Tlr9 in lung epithelium to induce rapid pathogen killing, and can be
154818	Tlr2	21482737	used as a therapeutic target to treat otherwise lethal pneumonia. Tlr2 is activated by gut commensal microbe, Bacteroides fragilis, to establish host-microbial
154818	Tlr2	21512004	symbiosis by promoting immunological tolerance.
			Tlr2 and Tnfsf11 signalling pathways are modulated by Porphromonas gingivalis to alter the differentiation states of osteoclasts resulting in bacteria-mediated bone loss.
154818	Tlr2	21566133	Tlr2 is expressed by Muller cells, principal glia of retina, and is responsible for generating robust bactericidal activity against Staphylococcus aureusand
154818	Tlr2	21602496	contributing to retinal innate defence.
154818	Tlr2	21698237	Tlr2 is required for rapid inflammasome activation in response to infection by cytosolic bacterial pathogens such as Francisella novicida.
			Tlr2-driven integration of inducible nitric oxide synthase (iNOS), Wnt-beta-Catenin and Notch1 signalling contributes to its capacity to regulate a battery of genes associated with T regulatory cell lineage commitment and towards modulation of
154818	Tlr2	21862586	defined set of effector functions in macrophages. Tlr2 directly recognizes glycogen, resulting in the activation of immunocytes such as macrophages to enhance the production of nitric oxide and
154818	Tlr2	21873606	inflammatory cytokines.

154818	Tlr2	22096480	Tlr2 and Tlr4 are crucial for in vivo recognition of Chlamydia pneumoniae. Tlr2/4 double-deficient mice were unable to control pneumonia caused by C. pneumoniae.
			Tlr2 signalling promotes protective vaccine-enhancing Th17 cell responses when cells are stimulated with early secreted antigenic target protein 6 (ESAT-6) expressed by the virulent Mycobacterium tuberculosis
154818	Tlr2	22102818	strain H37Rv but not by tuberculosis vaccine Bacillus Calmette-Guérin (BCG). Tlr2 recognizes Mycobacterium tuberculosis H37Rv cell surface lipoprotein MPT83, which induces the production of Tnf, Il6, and Il12b cytokines by
154818	Tlr2	22174456	macrophages and upregulates macrophage function. Mycobacterium abscessus glycopeptidolipid (GPL) prevents Tlr2-mediate induction of Il8 and Defb4a in
154818	Tlr2	22216191	respiratory epithelial cells. (Demonstrated in human) Tlr2 is expressed in the enteric nervous system (ENS) and intestinal smooth muscle layers. Its absence induces architectural and neurochemical coding changes in the ENS, leading to gut dysmotility and to
154818	Tlr2	23994200	higher inflammatory bowel diseases susceptibility Salmonella enterica serovar Typhimurium Î mmsbB that possesses a modified lipid A triggers exacerbated colitis in the absence of Nod1 and/or Nod2, which is
154818	Tlr2	25423082	likely due to increased Tlr2 stimulation.
154818	Tlr2	25505250	Adaptor proteins Ticam1 and Ticam2 have a novel function in Tlr2-mediated signal transduction. Retinoic acid treatment enhances Tlr2-dependent II10 production from T cells and this, in turn, potentiates T regulatory cell generation without the need for
154818	Tlr2	25826367	activation of antigen presenting cells. Proline-proline-glutamic acid 57 (PPE57), located on the mycobacterial cell surface, induces a T helper 1 immune response via Tlr2-mediated macrophage
154818	Tlr2	25586105	functions. Tlr2 is essential for the immune responses to cholera
154818	Tlr2	26078314	vaccination. Staphylococcal superantigen-like protein 3 (SSL3) interferes with Tlr2 activation at two stages. First by binding to Tlr2 and blocking ligand binding and second by interacting with an already formed Tlr2-
154818	Tlr2	26283364	lipopeptide complex, thus preventing TLR heterodimerization and downstream signalling. Peli3 is involved in endotoxin tolerance and functions as a negative regulator of Tlr2/4 signalling.
154818	Tlr2	26310831	Cytokine activation as a result of Tlr2 stimulation is
154818	Tlr2	26423153	mediated by the phagosomal trafficking molecule Ap3b1 (AP-3).

			Ap3b1 (AP-3), a lysosome-related organelle trafficking and biogenesis protein, is required for the production of pro-inflammatory cytokines in plasmacytoid dendritic cells upon recognition of viral nucleic acids by
173348	Ap3b1	21045126	endosomal Tlr7 or Tlr9. Ap3b1 is crucial for the trafficking of Tlr9 to specific endosomal compartments for the induction of type I interferon.
173348	Ap3b1	21119105	Cytokine activation as a result of Tlr2 stimulation is mediated by the phagosomal trafficking molecule
173348	Ap3b1	26423153	Ap3b1 (AP-3). Tlr9 binds CpG DNA and induces dendritic cell
195818	Tlr9	11286707	maturation in a MyD88-dependent manner in mice. Tlr9 engages in signalling crosstalk with Tlr4 during activation, and as a result, amplifies the inflammatory
195818	Tlr9	19923461	response of murine macrophages. Tlr9, 7, and 3 interact with the endoplasmic reticulum (ER) membrane protein Unc93b and this is essential for
195818	Tlr9	17452530	proper TLR signaling. Tlr9 signalling enhances the rate of acidification of the Salmonella-containing phagosome, and this acidification induces the expression of Salmonella pathogenecity genes that are necessary for intracellular survival, growth, and systemic infection. Tlr9
195818	Tlr9	21376231	deficiency rescues the high Salmonella susceptibility phenotype observed in Tlr2, Tlr4 double mutant mice. Tlr9 triggers plasmacytoid dendritic cells in Systemic Lupus Erythematosus patients upon recognition of self- antigens such as neutrophil extracellular traps (NETs).
195818	Tlr9	21389263	(Demonstrated in human) Tlr9 requires proteolytic processing in endolysosome by asparagine endopeptidase and cathepsin in the endolysosome to initiate signalling in response to
195818	Tlr9	21402738	DNA. Tlr9 deficiency reduced pancreatic edema,
195818	Tlr9	21439959	inflammation and pro-II1b expression in pacreatitis. Tlr9 synergistically interacts with Tlr2::Tlr6 in lung epithelium to induce rapid pathogen killing, and can be used as a therapeutic target to treat otherwise lethal
195818	Tlr9	21482737	neumonia. Tlr9 is proteolytically cleaved in the endosome to form a soluble Tlr9 (sTlr9), which inhibits Tlr9-dependent signalling and contributes to the prevention of
195818	Tlr9	21604257	autoimmune disease. Tlr9 activation is enhanced by increased levels of circulating histones, serving as a crucial link between initial damage and activation of innate immunity
195818	Tlr9	21721026	during sterile inflammation

			Tlr9 promotes macrophage Hifla levels, oxidative burst and nitric oxide production in response to group A Streptococcus (GAS), contributing to GAS clearance
			in vivo in both localized cutaneous and systemic
195818	Tlr9	21860217	infection models. Tlr9 is selectively compartmentalized to fungal
195818	Tlr9	21947771	<ul><li>phagosomes and negatively modulates macrophage anti-fungal effector functions.</li><li>TIr9 expression and signalling capacity oscillates with</li></ul>
195818	Tlr9	22342842	the circadian clock. Mitochondrial DNA that escapes from autophagy induces Tlr9 inflammatory responses in cardiomyocytes and is capable of inducing myocarditis
195818	Tlr9	22535248	and dilated cardiomyopathy. TLR9 activation by endogenous self-ligands generated
195818	Tlr9	23071157	during oxidative stress promotes platelet hyper- reactivity and thrombosis.
195818	Tlr9	23142781	Tlr9 contributes to the control of activated endogenous retroviruses (ERVs) and ERV-induced tumours. The N-terminal region of TLR9 is cleaved in the
195818	Tlr9	23752491	<ul><li>endolysosome to form an intracellular DNA sensor with the C-terminal TLR9.</li><li>Mir126-Kdr axis is an important regulator of the innate response. Mir126 controls the survival and function of</li></ul>
195818	Tlr9	24270517	plasmacytoid dendritic cells and regulates gene expression of Tlr7, Tlr9, Nfkb1 and Kdr.
195818	Tlr9	25600358	Dnase2a is required for Tlr9 activation by bacterial genomic DNA.
195818	Tlr9	26957214	Tyrosine phosphorylation is essential for Tlr9 protein stability and signalling. Tlr9 activation requires a single CpG positioned 4-6 nucleotides from the 5'-end and this activation is
195818	Tlr9	26416273	augmented by a 5'TCC sequence one to three nucleotides from the CG. Ifnb1 deficiency results in a partial suppression of the sterol pathway in macrophages during viral infections,
162198	Ifnb1	21408089	thereby linking the regulation of lipid metabolism pathway with interferon anti-viral defence responses. Ifnb1 secretion is greater upon viable E. coli infection in comparison to heat killed E. coli vaccine or LPS. The induction of Ifnb1 is dependent on Ticam1-Irf3 signalling.
162198	Ifnb1	21602824	Signaring.
162198	Ifnb1	22291574	Ifnb1 expression pattern during viral infection is a highly stochastic process influenced by cell-to-cell variability in viral induction processes. Ifnb1 production is fundamental to the efficient control of Listeria monocytogenes during the early innate phase of infection. NK cells treated with Ifnb1 during early infection were able to reduce bacterial titer in the
162198	Ifnb1	22912878	spleen and significantly improve survival of infected mice.

			The noncanonical NF ΰB pathway regulates histone modifications at the Ifnb1 promoter resulting in attenuated recruitment of Rela and histone demethylase, Kdm4a, to the Ifnb1 promoter. This
162198	Ifnbl	24656046	<ul><li>provides a mechanism for regulating the induction of type I interferons .</li><li>The innate immune system plays a role in immunogenic tumour recognition. Tumor-cell-derived DNA triggers Ifnb1 production and dendritic cell</li></ul>
162198	Ifnb1	25517615	activation via Tmem173 and Irf3 cytosolic DNA sensing pathways. Ifnb1 selectively restricts the transcriptional responses mediated by both the TLRs and the NOD-like receptors
162198	Ifnb1	26202980	<ul><li>in Salmonella enterica serovar Typhimurium infection in macrophages.</li><li>Atf3 plays an important role in modulating IFN responses in macrophages by controlling basal and</li></ul>
162198	Ifnb1	26416280	inducible levels of Ifnb1, as well as the expression of genes downstream of IFN signalling. Atf3 has been identified as a high-density lipoprotein
208712	Atf3	24317040	<ul> <li>â € inducible negative regulator of macrophage activation.</li> <li>Atf3 plays an important role in modulating IFN responses in macrophages by controlling basal and</li> </ul>
208712	Atf3	26416280	inducible levels of Ifnb1, as well as the expression of genes downstream of IFN signalling. NIrp3 mediated immune responses can be activated by RNA and mice lacking NIrp3, or other inflammasome components, exhibit increased mortality and a reduced immune response after influenza virus exposure.
178924	Nlrp3	19362020	Nlrp3(-/-) mice exhibit increased morbidity after infection with a pathogenic influenza A virus correlating with decreased neutrophil and monocyte
178924	Nlrp3	19362023	recruitment and reduced IL-1beta, IL-18, TNF-alpha, IL-6, KC, MIP2, and IFN-alpha production. Nlrp3 is directly activated by certain antibiotics and plays an important role in the antibiotic-mediated secretion of Il1b. In the case of polymyxin B, Nlrp3 was also required for the neutrophil influx into the
178924	Nlrp3	21278344	peritoneal cavity. Nlrp3 inflammasome is essential for host defence
178924	Nlrp3	21289120	against influenza and other RNA viruses (i.e. EMCV, VSV). Nlrp3 recruits adaptor protein Pycard and Casp1 to
178924	Nlrp3	21385879	form an Nlrp3 inflammasome complex in response to Varicella-Zoster Virus (VZV) infection. Nlrp3 is a component of the inflammasome and is
178924	Nlrp3	21439959	required for inflammation in acute pancreatitis.

			Nlrp3 is necessary to illicit Il1b response specific to viable, but not heat-killed, E. coli infections.
178924	Nlrp3	21602824	Nlrp3 inflammasome plays a role in innate immune responses against mucosal Candida infection. Nlrp3
178924	Nlrp3	22174673	limits the severity of infection when present in either the hematopoietic or stromal compartments. Nlrp3/Pycard inflammasome activation following human respiratory syncytial virus infection is dependent on the activation of Tlr2/Myd88/NF-kB and reactive oxygen species/potassium efflux.
178924	Nlrp3	22295065	(Demonstrated in human) MIR223 and EBV miR-BART15 regulate the NLRP3
178924	Nlrp3	22984081	inflammasome and IL-1beta production. Potassium efflux is a common trigger of NLRP3
178924	Nlrp3	23809161	Inflammasome activation by bacterial toxins and particulate matter Mitochondrial membrane potential is required for the
178924	Nlrp3	24127597	association of Nlrp3 and Mfn2. Mfns2 is required for the activation of Nlrp3 inflammasomes. 3, 4-methylenedioxy- $\hat{1}^2$ -nitrostyrene is a potent and
178924	Nlrp3	24265316	specific inhibitor of the NLRP3 inflammasome. Both Nlrp3 and Nlrp1a are important regulators of Toxoplasma proliferation and that IL18 signaling is
178924	Nlrp3	24549849	required to mediate host resistance to acute toxoplasmosis. Group B streptococcus induces II1b, and activates the NLRP3 inflammasome by a mechanism that requires hemolysin-mediated lysosomal leakage, which enhances the interaction of bacterial RNA with
178924	Nlrp3	24692555	NLRP3. Activation of the Nlrp3 inflammasome is detrimental during leishmaniasis. Mice lacking the inflammasome components Nlrp3, Pycard, Casp1 exhibit defective Il1b and Il18 production at the infection site and are resistant to cutaneous Leishmania major infection.
178924	Nlrp3	25689249	Nlrp3 is a novel molecular target for melatonin which requires Rora to blunt the NFkB/ NLRP3 connection
178924	Nlrp3	26045547	during sepsis. Uropathogenic Escherichia coli protein TcpC attenuates activation of the Nlrp3 inflammasome by
178924	Nlrp3	27214553	binding both Nlrp3 and Casp1. Nlrp3 deficiency protects mice from the development of type 1 diabetes by suppressing Th1 responses and impairing T-cell migration to pancreatic islets through the down-regulation of chemokine expression (Ccl5,
178924	Nlrp3	26305961	Cxcl10, Irf1) in islets. Nlrp3 and Casp2 are required for endoplasmic
178924	Nlrp3	26341399	reticulum stress-induced inflammation.

			Impairment of the mitochondrial electron transport chain by rotenone primes Nlrp3 inflammasome
178924	Nlrp3	26416893	activation upon costimulation with ATP. IRF3 is a transcription factor that activates type-1
63225	IRF3	9463386	interferon (IFN) and IFN responsive genes. IRF3 is directly activated after virus infection and functions as a key activator of the intermediate/early alpha/beta interferon (IFN) genes as well as the
63225	IRF3	11991981	RANTES chemokine gene. IRF3 transcription factor induces type I interferons (IFNs) and elicits innate antiviral response. TMEM173 (MITA) is a critical mediator of virus-triggered IRF3
63225	IRF3	18818105	activation and IFN expression. IRF3 phosphorylation is virus-inducible and results in IRF3 alteration of protein conformation to permit nuclear translocation, association with transcriptional
63225	IRF3	9566918	partners, and primary activation of interferon (IFN)- and IFN-responsive genes. IRF3 is strongly phosphorylated at the late stages of a
63225	IRF3	21768204	Sindbis virus infection to mount antiviral responses in human embryonic kidney cells. IRF3 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA and in the subsequent induction of type I IFNs. Mice lacking
63225	IRF3	21820332	Irf3/Irf7 are resistant to otherwise lethal cerebral malaria. (Demonstrated in mouse) IRF3 suppresses neuroinflammation through regulation of immunomodulatory MIR155 microRNA expression
63225	IRF3	22170100	in astrocytes. HIV accessory protein Vpu targets IRF3 to endolysosome for proteolytic degradation to avoid
63225	IRF3	22593165	antiviral immune responses. During the transcriptional response to Sendai virus
63225	IRF3	23994473	infection, POLR2F(RNA Pol II) is recruited by IRF3 and NFÎ <sup>o</sup> B to control virus induced gene activation. Vpu, an accessory protein encoded by HIV-1, contributes to the attenuation of the anti-viral response by partial inactivation of IRF3 while host cells undergo
63225	IRF3	25352594	apoptosis. Hepatitis B virus (HBV) polymerase inhibits TMEM173-stimulated IRF3 activation and IFNB1
63225	IRF3	25505063	induction. Stimulation of TMEM173-dependent IRF3 activation by ultraviolet radiation is due to apoptotic signalling- dependent disruption of ULK1, a pro-autophagic
63225	IRF3	25792739	protein that negatively regulates TMEM173. PQBP1 directly binds to reverse-transcribed HIV-1 DNA and interacts with MB21D1 to initiate an IRF3-
63225	IRF3	26046437	dependent innate response

			4-(2-chloro-6-fluorobenzyl)-N-(furan-2-ylmethyl)-3-
			oxo-3,4-dihydro-2H-benzo[b][1,4]thiazine-6-
			carboxamide (G10) requires STING to trigger
			IRF3/IFN-associated transcription in human fibroblasts
			and subsequently blocking replication of Chikungunya virus, Venezuelan Encephalitis virus, and Sindbis
63225	IRF3	26646986	virus, venezueran Encephantis virus, and sindois virus.
05220	nu s	20010/00	Influenza B virus induces IRF3 activation and IL29
			(IFNL1) gene expression without a requirement for
			viral protein synthesis or replication and DDX58 is the
(2225		2(2701(0	critical pattern recognition receptor needed for IRF3
63225	IRF3	26378160	activation. IL29 is able to efficiently inhibit Herpes Simplex Virus
			Type-1 replication in neuronal cells by inducing the
			expression of TLR as well as activate the TLR-
50119	IL29	21499846	mediated interferon antiviral pathway.
			IFNL1 (IL29), IFNL2 and IFNL3 have different effects
50110	11.20	24041(72	on Toll-like receptor-related gene expression in HepG2 cells.
50119	IL29	24041672	IFN- $\hat{I} \gg 1$ is able to augment TLR-mediated B cell
			activation, partially attributed to an upregulation of
50119	IL29	26130701	TLR7 expression
			Influenza B virus induces IRF3 activation and IL29
			(IFNL1) gene expression without a requirement for
			viral protein synthesis or replication and DDX58 is the critical pattern recognition receptor needed for IRF3
50119	IL29	26378160	activation.
00119		200,0100	TRAF6 is an adapter protein linking kinases to TNF
			receptor, and IL-1 receptor signalling pathways, and
40102	TRAF6	16184196	also has E3 ubiquitin ligase activity.
			TRAF6 autoubiquitination and its interaction with Ubc13 are dependent on zinc finger 1 (ZF1) motif and
			an intact RING domain, necessary elements in
40102	TRAF6	18617513	signalling by IL-1, LPS and RANKL.
			TRAF6 is specifically required for the Smad-
			independent activation of JNK and p38, and its
			carboxyl TRAF homology domain physically interacts
			with TGF-beta receptors that activate JNK and p38 through a mechanism similar to that operating in the
40102	TRAF6	18922473	interleukin-1beta/Toll-like receptor pathway.
			TRAF6 and MEK kinase 1 (MEKK1) play a pivotal
			role in the retinoic-acid-inducible gene-I (RIG-I)-like
			helicase antiviral pathway, where TRAF6 and MEKK1
			activate NF-kappaB and mitogen-activated protein
40102	TRAF6	18984593	kinases via MAVS and this is critical for the optimal induction of type I interferons.
10102		10/010/0	TRAF6 negatively regulates TNFalpha-induced NF-
40102	TRAF6	19091594	kappaB activation through its ubiquitin ligase activity.
			TRAF6 regulates several signalling cascades in innate
40102	TRAF6	12140561	immunity, adaptive immunity and bone homeostasis.

40102	TRAF6	20449947	TRAF6 competes with TRAF2 for CD40 binding to regulate NF-kappaB activation in human B lymphocytes, thereby limiting the capacity of CD40 engagement to induce NF-kappaB activation. TRAF6 is autoinhibited by an intramolecular interaction which is counteracted by trans- ubiquitination and TRAF auto-ubiquitination is a
40102	TRAF6	20512936	means of sustaining an open conformation active in downstream signalling. TRAF6 interacts with CSF2RB to mediate NF-kappaB signalling, demonstrating a novel TRAF6-dependent signalling pathway association with a type I cytokine
40102	TRAF6	20622119	receptor. TRAF6 is a E3 ubiquitin ligase that activates NFKB pathway in response to innate and adaptive immunity stimuli. TRAF6 protein contains a highly conserved TRAF-C domain that contributes to oligomerization and its interaction to upstream signalling molecules and a RING domain dimerization interface that is functionally important for ubiquitination and the activation of NFKB.
40102	TRAF6	21185369	TRAF6 is polyubiquitinated and disassembled during endotoxin tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of anti- inflammatory or anti-microbial mediators.
40102	TRAF6	21220427	
40102	TRAF6	22033459	TRAF6 is degraded in the proteasome upon TLR stimulation in macrophages. (Demonstrated in mice) Following NOD2 activation, IRF4 interacts with MYD88, TRAF6, and RIPK2 and downregulates K63- linked polyubiquitinylation of RICK and TRAF6
40102	TRAF6	24670424	leading to disruption of NFkB activation pathways. MIR146A is a potent negative regulator of the innate immune response in keratinocytes through
40102	TRAF6	24670381	downregulation of the IRAK1/TRAF6/NFÎ <sup>o</sup> B pathway. STAT1 is directly recruited to TRAF6, demonstrating cross-talk between the TLR and JAK/STAT signalling pathways, and this direct activation of STAT1 by TLR signalling suggests a crucial role for STAT1 in TLR-
40102	TRAF6	25027037	induced inflammation. Demonstrated in mice. ECSIT binds to MAP3K7 and TRAF6 to form a
40102	TRAF6	25371197	complex that plays a pivotal role in activating TLR4- mediated NF-kB signalling. Reversible arginine methylation of TRAF6 is regulated
40102	TRAF6	26221041	by PRMT1 and JMJD6 and this in turn regulates TRAF6-dependent TLR signalling.
40102	TRAF6	26385923	MAVS directly interacts with TRAF6 through its potential TRAF6-binding motif 2.

49080	MAVS	19036819	MAVS oligomer is essential in the formation of a multiprotein membrane-associated signalling complex that enables downstream activation of IRF3 and NF- kappaB in antiviral innate immunity. MAVS is a caspase recruitment domain (CARD)- containing adaptor protein that interacts with DDX58 (RIG-I) and recruits CHUK (IKKalpha), IKBKB
49080	MAVS	16177806	<ul> <li>(IKKbeta) and IKBKE (IKKepsilon) kinases by means of its C-terminal region, leading to the activation of NF-kappaB and IRF3.</li> <li>MAVS is an adaptor protein that contains an N-terminal caspase recruitment domain (CARD)-like domain and a C-terminal transmembrane domain, both of which are essential for MAVS signalling, while the</li> </ul>
49080	MAVS	16125763	transmembrane domain also targets MAVS to the mitochondria. MAVS facilitates cell death by disrupting the mitochondrial membrane potential and by activating caspases.
49080	MAVS	20032188	MAVS-dependent RIG-I-like receptor (RLR)
49080	MAVS	20140199	signalling regulates the quantity, quality, and balance of the immune response to West Nile virus (WNV) infection. MAVS in peroxisomes induces a rapid interferon- independent expression of defence factors that provide short-term protection, whereas mitochondrial MAVS
			activates an interferon-dependent signalling pathway with delayed kinetics, which amplifies and stabilizes
49080	MAVS	20451243	the antiviral response. MAVS interacts with hepatitis B virus X protein and
49080	MAVS	20554965	this promotes the degradation of MAVS via Lys(136) ubiquitination, preventing the induction of IFN-beta. MAVS (IPS-1) interacts with MFN1 upon virus- infection or 5'ppp-RNA activation through
49080	MAVS	20661427	redistribution of MAVS to form speck-like aggregates in cells. MAVS (IPS1) plays an important role in regulating the
49080	MAVS	20699220	host anti-viral response by binding to viral polymerase and inhibiting IFN-beta production. MAVS negatively regulates the stability of voltage- dependent anion channel 1 (VDAC1) and thereby
49080	MAVS	21110072	inhibits apoptosis in the response to release of cytochrome c. The MAVS signalling pathway in non-myeloid cells is
49080	MAVS	21454965	crucial for dsRNA-mediated natural killer cell activation. (Demonstrated in murine model) Tyrosine phosphorylation of MAVS at amino acid
49080	MAVS	22844514	residue Tyr9 is critical for the induction of IFNB signalling.

			MAVS mRNA is degraded in response to foreign RNA
49080	MAVS	23028806	and poly(I:C) to suppress hyper-immune reaction in late-phase antiviral signalling. MAVS is required for optimal NLRP3 inflammasome
			activity by mediating mitochondrial recruitment of
49080	MAVS	23582325	NLRP3.
49080	MAVS	23555247	MAVS is targeted by enterovirus protease to evade antiviral immunity.
			The binding of MAVS to Traf2, Traf5, and Traf6 is dependent on virus infection and MAVS polymerization. The TRAF proteins promote
49080	MAVS	23951545	ubiquitination that recruits IKBKG binding to the MAVS signalling complex. Upon viral infection, MAVS recruits MKK7 onto
			mitochondria, leading to the induction of apoptosis by
49080	MAVS	24651600	MAP2K7 activated MAPK9
			Macrocyclic NS3-4A resistance-associated amino acid
			variants (RAVs) with substitutions at residue D168 of the hepatitis C virus protease result in an increased
			capacity of NS3-4A to cleave MAVS and suppress
49080	MAVS	25463536	IFNB1 induction.
			RNA cleavage products, catalyzed by RNASEL, bind to DHX33 to facilitate the formation of a complex with
49080	MAVS	25816776	MAVS and NLRP3 during viral infection.
			HACE1 plays an inhibitory role in virus-induced
49080	MAVS	27213432	signalling by disrupting the MAVS-TRAF3 complex.
10000	MAVS	26246171	MARCH5 modulates MAVS-mediated antiviral
49080	MAV5	26246171	signalling, preventing excessive immune reactions. MAVS directly interacts with TRAF6 through its
49080	MAVS	26385923	potential TRAF6-binding motif 2.
			IKBKB regulates late-phase allergic reactions
10007	WDVD	10(00471	promoted by the release of pro-inflammatory cytokines
19987	IKBKB	18692471	in an NF-kappaB-dependent manner. Phosphorylated IKBKB is conjugated with a
			monoubiquitin by the E3 ubiquitin ligase TRIM21
			leading to down-regulatioin of IKBKB-induced NF-
			kappaB signalling. The TRIM21-mediated
19987	IKBKB	20627395	monoubiquitination is involved in the translocation of active IKBKB to autophagosomes.
17707	inditid	20027575	IKBKB and other IKK kinases regulate each other by
			an intricate network involving phosphorylation of their
			catalytic and regulatory (NEMO, TANK) subunits to
19987	IKBKB	21138416	balance their activities during innate immunity.
17701			GNB2L1 (RACK1) negatively regulates NF Î ° B
			activation by interacting with CHUK and IKBKB. The
19987	IKBKB	24323043	interaction interferes with the recruitment of the IKK complex to TRAF2.
1770/	INDND	24323043	The reversible ubiquitin editing of NLRC5 determines
			NLRC5†IKBKB interaction dynamics and plays a
19987	IKBKB	26620909	crucial role in precisely regulating NFήB signalling

			Enterovirus 71 2C protein binds to RELA and IKBKB to inhibit NF-kB activation and evade innate immune
19987	IKBKB	26394554	defenses. RELA, NF-kappaB p65 subunit, is involved in the transcription regulation of many genes including those genes involved in apoptosis, response to stress and
57543	RELA	11980335	inflammation. RELA is a subunit of NFKB and is not essential for virus-stimulated IFNB expression, instead, RELA sustains autocrine IFNB signalling prior to infection. The absence of RELA causes significant delays in IFNB induction and consequently defective secondary antiviral gene expression. RELA maintains autocrine IFNB signalling in uninfected cells, facilitates inflammatory and adaptive immune responses following infection, and promotes infected cell survival during this process.
57543	RELA	21209118 21216972	RELA is critical for pulmonary host defence during Streptococcus pneumoniae pneumonia in alveolar macrophages. During pneumococcal pneumonia, only the earliest induction of cytokines depends on transcription regulated by RELA in myeloid cells, and this transcriptional activity contributes to effective immunity. (Demonstrated in murine model)
57543	KELA	21216972	RELA is required for IL17A production in T cell in response to bacterial infection. RELA deficient T cells resulted in a diminished innate immune response to E.
57543	RELA	21419662	coli infection. (Demonstrated in murine model) A RELA isoform, p43, lacks the transactivation domain but is still able to potentiate anti-viral innate
57543	RELA	23271966	immunity. During the transcriptional response to Sendai virus infection, POLR2F(RNA Pol II) is recruited by IRF3
57543	RELA	23994473	and NF PB to control virus induced gene activation. Paramyxoviruses trigger the DNA-damage response, a pathway required for RPS6KA5 activation of phospho Ser 276 RELA formation to trigger the IRF7-DDX58 amplification loop necessary for mucosal interferon
57543	RELA	25520509	production. Human papillomaviruses impair the acetylation of NFΰB/RelA K310 in keratinocytes by augmenting the
57543	RELA	26055519	expression of interferon-related developmental regulator 1 (IFRD1) in an EGFR-dependent manner. Haploinsufficiency of A20 (HA20) is caused by high- penetrance loss-of-function germline mutations in TNFAIP3 with increased degradation of NFKBIA, nuclear translocation of RELA, increased expression of NF Î ° B mediated proinflammatory cytokines, and
57543	RELA	26642243	defective deubiquitinating activity.

			MIR223 regulates macrophage function by modulating cytokine production and NF-ΰB activation through
57543	RELA	26296289	inhibition of RELA phosphorylation and nuclear translocation. Enterovirus 71 2C protein binds to RELA and IKBKB
57543	RELA	26394554	to inhibit NF-kB activation and evade innate immune defenses. Irf3 is strongly phosphorylated at the late stages of a Sindbis virus infection to mount antiviral responses in
181123	Irf3	21768204	human embryonic kidney cells. (Demonstrated in human) Irf3 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA and in the subsequent induction of type I IFNs. Mice lacking
181123	Irf3	21820332	Irf3/Irf7 are resistant to otherwise lethal cerebral malaria. Irf3 suppresses neuroinflammation through regulation
181123	Irf3	22170100	of immunomodulatory mmu-mir-155 microRNA expression in astrocytes. (Demonstrated in human) HIV accessory protein Vpu targets Irf3 to
181123	Irf3	22593165	endolysosome for proteolytic degradation to avoid antiviral immune responses. (Demonstrated in human)
181123	Irf3	23028052	Endoplasmic reticulum stress primes macrophages to respond to innate immunity stimuli by activating IRF3. Plasmodium RNA is a pathogen-associated molecular pattern (PAMP) capable of activating a type I IFN response via the cytosolic pattern recognition receptors
181123	Irf3	24362933	Ifih1 and Mavs, as well as via transcription factors Irf3 and Irf7. Il28ra (Ifnlr1), Stat1 and Irf3 are required for
181123	Irf3	25431490	antibiotics to prevent persistent murine norovirus infection. The innate immune system plays a role in immunogenic tumour recognition. Tumor-cell-derived DNA triggers Ifnb1 production and dendritic cell
181123	Irf3	25517615	activation via Tmem173 and Irf3 cytosolic DNA sensing pathways. Aberrant mitochondrial DNA (mtDNA) packaging promotes escape of mtDNA into the cytosol, where it engages the DNA sensor Mb21d1 and promotes Tmem173-Irf3-dependent signalling to elevate IFN-
181123	Irf3	25642965	stimulated gene expression, potentiate type I IFN responses and confer broad viral resistance. Ppp4c, a serine/threonine phosphatase, directly binds to Tbk1 upon virus infection to dephosphorylate Tbk1 and inhibit Tbk1 activation, and subsequently restrain
181123	Irf3	26363053	Irf3 activation, resulting in suppressed production of type I IFN and IFN-stimulated genes. Tbk1 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA and in the
192882	Tbk1	21820332	subsequent induction of type I IFNs. Mice lacking Tbk1 are resistant to otherwise lethal cerebral malaria.

192882	Tbk1	22921120	Tbk1 is a key regulator of immunological autophagy and is responsible for autophagosome maturation into bactericidal organelles.
			Ppp4c, a serine/threonine phosphatase, directly binds to Tbk1 upon virus infection to dephosphorylate Tbk1 and inhibit Tbk1 activation, and subsequently restrain
192882	Tbk1	26363053	Irf3 activation, resulting in suppressed production of type I IFN and IFN-stimulated genes. Ppp4c, a serine/threonine phosphatase, directly binds to
200020	D (	2 ( 2 ( 2 0 5 2	Tbk1 upon virus infection to dephosphorylate Tbk1 and inhibit Tbk1 activation, and subsequently restrain Irf3 activation, resulting in suppressed production of
209838	Ppp4c	26363053	type I IFN and IFN-stimulated genes. Myd88 restricts West Nile virus (WNV) by inhibiting replication in subsets of cells and modulating expression of chemokines that regulate immune cell
204565	Myd88	20881045	migration into the central nervous system. Myd88 signalling plays an important role for resisting primary influenza virus infection but is dispensable for
204565	Myd88	20943980	protection against a secondary lethal challenge. Myd88 is essential in restricting Tlr3 signaling and the host protection from unwanted immunopathologies associated with excessive production of Ifnb1. Myd88 inhibits Tlr3 signalling by impairing Ikbke-mediated
			induction of Irf3, and consequently the expression of Ifnb1 and Ccl5.
204565	Myd88	21248248	Myd88 is activated by MHC class II in response to staphylococcal enterotoxins and is crucial for the induction of pro-inflammatory cytokines.
204565	Myd88	21283748	
			MYD88 is a key signalling adapter in TLR signalling. MYD88 aggregates in the cell as distinct foci and co- localizes with IRAK4 in these Myddosomes - the
204565	Myd88	21325272	formation of which is required for MYD88 function. (Demonstrated in human) Myd88 is required in dendritic cells stimulated with Tlr9 ligand for the enhancement of T cell-dependent
204565	Myd88	21353603	antibody response. In addition, Myd88 is required in B cells to facilitate strong anti-viral antibody responses. Myd88 deficient macrophages displayed impaired interaction with fungal yeast cells and produced low levels of pro-inflammatory cytokines. Myd88 signalling is important in the activation of fungicidal mechanisms and the induction of protective innate
204565	Myd88	21422180	immune responses against P. brasiliensis. Myd88 mediates cytoskeletal remodelling and late spreading of lipopolysaccharide (LPS)-stimulated
204565	Myd88	22028692	macrophages.

			Myd88-dependent recruitment of inflammatory monocytes and dendritic cells to the lungs are key
			initial cellular responses required for early protection
204565	Myd88	22025508	from Burkholderia mallei infection.
			Myd88 deficiency results in delayed recruitment of phagocytes and defective production of
			proinflammatory cytokines in response to Salmonella
204565	Myd88	22386951	infection.
204565	<b>M</b> 100	22401177	Myd88 signalling in intestinal epithelial cells is crucial
204565	Myd88	22491177	for the maintenance of gut microbiota homeostasis. Myd88 mediated production of reactive oxygen species
			(ROS) is essential for the induction of Il12 by lactic
204565	Myd88	22536449	acid bacteria.
			Flagellin-specific IgG1 antibody response is induced
204565	Myd88	24442437	through a Tlr5-, inflammasome-, and Myd88- independent pathway.
201000	Wydoo	21112137	The SF3A/SF3B mRNA splicing complexes regulate
			the innate immune response in part by regulating
204565	M 100	24204290	Myd88S levels, which modulate the extent of the
204565	Myd88	24204290	innate immune response through Tlr4. Mir149 negatively regulates TLR/Myd88 mediated
			inflammatory responses in macrophages by targeting
204565	Myd88	24375488	Myd88 mRNA.
204565	N 100	25440706	Myd88 and Ticam1 pathways differently regulate Tlr4-
204565	Myd88	25448706	induced immune responses in B cells. Ticam1 but not Myd88 signalling is critical for the Trl4
			protective adjuvant effect in neonates; where Ticam1(-
			/-) but not Myd88(-/-) neonates are highly susceptible
204565	Myd88	25548220	to Escherichia coli peritonitis and bacteremia.
			Treml4 is an essential positive regulator of Tlr7 signalling. Treml4(-/-) macrophages are
			hyporesponsive to Tlr7 agonists and fail to produce
			type I interferons due to impaired phosphorylation of
004565	N 100	05040064	Stat1 by Mapk14 and decreased recruitment of Myd88
204565	Myd88	25848864	to Tlr7. Intracellular Sef/IL-17R (SEFIR) domain of Il17rd
			targets TIR adaptor proteins Myd88, Tirap, Ticam1,
			Ticam2 and Traf6 to inhibit TLR downstream
204565	Myd88	25808990	signalling.
			Map1s (Mtap1s) controls bacterial phagocytosis through TLR signalling by interacting directly with
204565	Myd88	26565030	Myd88.
	2		Hepatocyte Myd88 affects bile acids, gut microbiota
004565	N 100	07106570	and metabolome contributing to regulation of glucose
204565	Myd88	27196572	and lipid metabolism. Extracellular RNA of cardiac origin exhibits a potent
			pro-inflammatory property in vitro and in vivo and
			induces cytokine production through Tlr7-Myd88
204565	Myd88	26363072	signalling.
			Tlr7, 9, and 3 interact with the endoplasmic reticulum (ER) membrane protein UNC93B and this is essential
185532	Tlr7	17452530	for proper TLR signaling.

			Tlr7 is expressed in C-fiber primary sensory neurons and is important for inducing itch (pruritus), but is not necessary for eliciting mechanical, thermal, inflammatory and neuropathic pain in mice
185532	Tlr7	21037581	Tlr7 signaling pathway plays a pivotal role in fungal pathogen recognition and is essential for the
185532	Tlr7	21282509	subsequent Ifnb signaling.
185532	Tlr7	21402738	Tlr7 requires proteolytic processing in endolysosome by asparagine endopeptidase and cathepsin in the endolysosome to initiate signalling.
			Tlr7 agonist, such as imidazoquinolines, accumulate in the MHC class II loading compartment - this pH-
185532	Tlr7	21487111	dependent localization is required for the activation of plasmacytoid dendritic cells. (Demonstrated in human) TLR7 inflammatory signalling leads to cardiac fibrosis
185532	Tlr7	21730058	in autoimmune associated congenital heart block (Demonstrated in human)
			Tlr7 and Tlr8 are translocated from the endoplasmic reticulum to the endosome in the presence of antiphospholipid antibodies, as a consequence, plasmacytoid dendritic cells become dramatically sensitized to Tlr7/8 agonists and this may play a role in
185532	Tlr7	21734241	systemic autoimmunity. Tlr7 is responsible for the detection of retroviruses and serves as a key checkpoint controlling the development
185532	Tlr7	21998589	of germinal center B cells. Tlr7 signalling induces autophagy in HIV-infected
185532	Tlr7	22396599	plasmacytoid dendritic cells; this process is necessary for the induction of IFN-alpha. (Demonstrated in mice) Tlr7 binds to exosomal Mir21 and Mir29a secreted by
185532	Tlr7	22753494	tumour cells and initiates a prometastatic inflammatory response. Aberrant TLR7 activation induces Epstein-Barr viral
185532	Tlr7	22952664	protein LMP1 expression, which exacerbates IFN production in lupus patients. (Demonstrated in human)
185532	Tlr7	23142781	<ul><li>Tlr7 contributes to the control of activated endogenous retroviruses (ERVs) and ERV-induced tumours.</li><li>Mir126-Kdr axis is an important regulator of the innate response. Mir126 controls the survival and function of</li></ul>
185532	Tlr7	24270517	plasmacytoid dendritic cells and regulates gene expression of Tlr7, Tlr9, Nfkb1 and Kdr. Treml4 is an essential positive regulator of Tlr7 signalling. Treml4(-/-) macrophages are hyporesponsive to Tlr7 agonists and fail to produce type I interferons due to impaired phosphorylation of Stat1 by Mapk14 and decreased recruitment of Myd88
185532	Tlr7	25848864	to Tlr7.

			Neuronal Tlr7 recognizes endogenous ligands such as the miRNAs Let7c and miR21 and plays a negative role in controlling neuronal growth in a cell-
185532	Tlr7	25917529	autonomous manner. Extracellular RNA of cardiac origin exhibits a potent pro-inflammatory property in vitro and in vivo and
185532	Tlr7	26363072	induces cytokine production through Tlr7-Myd88 signalling. Epithelial cell-intrinsic Chuk expression and Tslp
134008	Tslp	26371187	regulate group 3 innate lymphoid cell responses required to maintain intestinal barrier immunity. Chuk has a key role in the negative feedback of NF-kB canonical signalling by orchestrating the assembly of the A20 ubiquitin-editing complex to limit
169132	Chuk	21765415	inflammatory gene activation in response to proinflammatory stimuli such as Tnf and II1. Epithelial cell-intrinsic Chuk expression and Tslp
169132	Chuk	26371187	regulate group 3 innate lymphoid cell responses required to maintain intestinal barrier immunity. IFIT5 is a positive regulator in IKK phosphorylation
81992	IFIT5	26334375	and NF-ΰB activation. Transcriptional activation of Adar by IFN occurs in the
197036	Stat2	26335850	absence of Stat1 by a non-canonical Stat2-dependent pathway. Stat1 phosphorylation at Ser708 is a key event in the
153162	Stat1	22065572	IFN signalling pathway that imparts anti-viral immunity to restrict West Nile virus infection.
153162	Stat1	22425562	Histone deacetylase inhibitors prevent Ifng-mediated phosphorylation of Stat1. (Demonstrated in human) Il28ra (Ifnlr1), Stat1 and Irf3 are required for
153162	Stat1	25431490	antibiotics to prevent persistent murine norovirus infection. Cd81 inhibits Rac1/Stat1 activation and negatively
153162	Stat1	25972472	regulates the defence mechanisms to Listeria monocytogenes infection. Treml4 is an essential positive regulator of Tlr7 signalling. Treml4(-/-) macrophages are hyporesponsive to Tlr7 agonists and fail to produce type I interferons due to impaired phosphorylation of
153162	Stat1	25848864	Stat1 by Mapk14 and decreased recruitment of Myd88 to Tlr7. Transcriptional activation of Adar by IFN occurs in the
153162	Stat1	26335850	absence of Stat1 by a non-canonical Stat2-dependent pathway. Adar destabalizes RNA structure by the deamination of
164791	Adar	21809195	adenosine to inosine, and therefore is able to disrupt replication of dsRNA viruses in the host. Transcriptional activation of Adar by IFN occurs in the
164791	Adar	26335850	absence of Stat1 by a non-canonical Stat2-dependent pathway.

			The Ticam1 signalling pathway in murine dendritic cells is crucial for dsRNA-mediated natural killer cell
191912	Ticam1	21454965	activation.
191912	Ticam1	21494017	Ticam1 deficiency results in the impairment of LPS- stimulated TNF-alpha protein translation. Ticam1 is crucial for Nlrp3 inflammasome activation in response specific to viable, but not heat-killed, E. coli infections.
191912	Ticam1	21602824	Ticam1 is proteolytically cleaved by Enterovirus 71 to inhibit the induction of innate immunity by Tlr3- signalling. Ticam1 cleavage results in the inhibition of NFkB and IFNB promoter activation. (Demonstrated in
191912	Ticam1	21697485	human) Ticam1 forms a dsRNA sensor complex with components Ddx1, Ddx21 and Dhx36 to trigger the type I interferon and cytokine response to poly I:C,
191912	Ticam1	21703541	influenza A virus, and reovirus. Ticam1 is a potent negative regulator of TLR agonist-
191912	Ticam1	21760953	triggered immune responses, specifically suppressing Il12 in dendritic cells and Ifng in natural killer cells. Ticam1-Tlr3-mediated signalling pathway plays an essential role in the anti-viral response against
191912	Ticam1	22072781	essential fole in the anti-vital response against poliovirus infection. Ticam1 plays a role in host resistance to Gram-negative enteropathogens. Ticam1-mediated protective immunity is orchestrated by macrophage-induced IFN-
191912	Ticam1	22124111	beta and natural killer cell production of IFN-gamma. Ticam1 forms a complex with Ripk3 upon Toll-like receptors (TLR) 3 and 4 activation resulting in Ripk3- dependent but TNF-independent necrosis in
191912	Ticam1	22123964	macrophages. High-potency Tlr4 agonists can act as clinically useful vaccine adjuvants by selectively activating Ticam1- dependent immunostimulatory signalling events and only weakly activating potentially harmful Myd88-
191912	Ticam1	25389373	dependent inflammatory responses. Myd88 and Ticam1 pathways differently regulate Tlr4-
191912	Ticam1	25448706	induced immune responses in B cells. Adaptor proteins Ticam1 and Ticam2 have a novel
191912	Ticam1	25505250	function in Tlr2-mediated signal transduction. Ticam1 but not Myd88 signalling is critical for the Trl4 protective adjuvant effect in neonates; where Ticam1(- /-) but not Myd88(-/-) neonates are highly susceptible
191912	Ticam1	25548220	to Escherichia coli peritonitis and bacteremia. Wdfy1 is a crucial adaptor protein in the Tlr3/4 signalling pathway. Wdfy1 interacts with Tlr3 and Tlr4 and mediates the recruitment of Ticam1 to these
191912	Ticam1	25736436	receptors.

191912 191912	Ticam1 Ticam1	25808990 26651944	Intracellular Sef/IL-17R (SEFIR) domain of Il17rd targets TIR adaptor proteins Myd88, Tirap, Ticam1, Ticam2 and Traf6 to inhibit TLR downstream signalling. Yersinia pseudotuberculosis type III secretion system effector, YopJ, suppresses Trif(Ticam1)-dependent responses during infection of primary phagocytic cells, including dendritic cells and macrophages. Ticam1-dependent type I interferon signalling in T cells is essential to Th1 lineage differentiation and reactivation of memory T cells. Ticam1 activated memory T cells facilitate local neutrophil influx and
			enhance bacterial elimination.
191912	Ticam1	26351279	CAMP represents a potent antimicrobial and cell- stimulating agent, most abundantly expressed in peripheral organs such as lung and skin during
32341	CAMP	19625657	inflammation.
32341	САМР	19748465	CAMP, a protein that has direct antimicrobial activity, serves as a mediator of vitamin D3-induced autophagy. CAMP (LL-37) modulates IFN-gamma responses during both the innate and adaptive phases of immune
32341	САМР	19812202	responses, indicating an immunomodulatory role for this endogenous peptide. CAMP has both antimicrobial and regenerative capabilities and promotes high glucose-attenuated epithelial wound healing via EGFR transactivation in organ cultured corneas.
32341	CAMP	19797203	CAMP is involved in various aspects of skin biology,
32341	САМР	18923446	<ul><li>including protection against infection, wound healing,</li><li>and also in psoriasis where it suppresses apoptosis in keratinocytes.</li><li>CAMP induces endothelium-dependent relaxation in human omental veins, or vasodilation, mediated via an</li></ul>
32341	CAMP	18397922	effect on endothelial ALX. CAMP enhances delivery of CpG oligodeoxynucleotides to stimulate immune cells and this is independent of its amphipathic structure and its
32341	CAMP	20042575	bactericidal property. CAMP has dual function as an antimicrobial agent
32341	САМР	20036634	against bacterial target cells and a cell penetrating peptide that can deliver nucleic acids into the host cells. CAMP decreases collagen expression at mRNA and protein levels in human dermal fibroblasts (HDFs) and this inhibition is dependent on phosphorylation of extracellular signal-regulated kinase (ERK). Vitamin C attenuates ERK signalling to inhibit the regulation of
32341	CAMP	20163451	collagen production by CAMP in HDFs.

32341	САМР	20190140	CAMP and human beta-defensins (hBDs) antimicrobial peptides induce the secretion of a pruritogenic cytokine IL-31 by human mast cells. CAMP (LL37) directs macrophage differentiation toward macrophages with a pro-inflammatory signature
32341	САМР	20610648	and this requires internalization of the peptide, resulting in low production of IL-10 and profound production of IL-12p40 upon LPS stimulation. CAMP (LL37) converts self-RNA into a trigger of TLR7 and TLR8 in human dendritic cells (DC),
32341	CAMP	19703986	leading to production of TNF-alpha and IL6 and the differentiation of myeloid DCs into mature DCs. CAMP (LL-37) attenuates lethal sepsis/endotoxin shock by suppressing the LPS-induced apoptosis of vascular and hepatic endothelial cells. LL-37 was found
32341	САМР	21393634	to inhibit the binding of LPS to the LPS receptors expressed on the cells. CAMP (LL37) is found in high concentrations within neutrophil extracellular traps (NETs). CAMP is a neutrophil protein that facilitates the uptake and
32341	САМР	21389264	recognition of mammalian DNA by plasmacytoid dendritic cells, and may play a role in Systemic Lupus Erthermatosus autoimmunity. CAMP (LL-37) dramatically reduced TNFA and nitric oxide levels produced by LPS and IFNG-polarized M1 macrophages, in addition LL-37-treated M1
32341	САМР	21441450	<ul> <li>macrophages were more efficient at suppressing tumour growth in vitro. This demonstrates the selective ability of LL-37 to decrease production of LPS-induced pro-inflammatory cytokines in macrophages, while leaving other crucial anti-inflammatory M1 and M2 macrophage functions unaltered.</li> <li>CAMP (LL-37), at sufficiently low concentrations, is able to reduce fungal infectivity by inhibiting C. albicans adhesion to plastic surfaces, oral epidermoid cells, and the urinary bladders of female mice. The inhibitory effects of LL-37 on cell adhesion and</li> </ul>
32341	CAMP	21448240	aggregation were mediated by its preferential binding to mannan and chitin in the fungal cell wall. CAMP (LL-37) translocates across the E. coli outer
32341	CAMP	21464330	membrane and halts bacterial growth by interfering cell wall biogenesis. CAMP (LL-37) confers protective immunity against psoriasis by neutralizing cytosolic DNA in keratinocytes and blocking the formation of AIM2 inflammasomes.
32341	CAMP	21562230	CAMP protects against colitis induction in mice. The increased expression of CAMP in monocytes involves
32341	CAMP	21762664	the activation of TLR9/ERK signalling pathway by bacterial DNA. (Demonstrated in mouse)

			CAMP expression is induced upon endoplasmic
32341	CAMP	21832078	reticulum stress via NF-kB-C/EBP-alpha activation.
32341	CAMP	22031815	LL-37 (CAMP) reduces influenza A viral load and
			disease severity in mice.
32341	CAMP	23328115	CAMP (LL-37) is downregulated during septic shock. (S)-methyl 2-(hexanamide)-3-(4-hydroxyphenyl)
			propanoate (MHP) activates SPHK1 to stimulate
			CAMP production and enhance epidermal
32341	CAMP	26113114	antimicrobial defence.
			Cleavage of CAMP by cathepsins CTSS and CTSK
			impairs its antimicrobial activity against Pseudomonas
32341	CAMP	25884905	aeruginosa and Staphylococcus aureus.
22241	CAMD	26979966	Carbamylation of CAMP affects its bactericidal,
32341	CAMP	26878866	cytotoxic and immunomodulatory function. CAMP modulates the response of macrophages during
			mycobacterial infection controlling the expression of
32341	CAMP	26351280	pro-inflammatory and anti-inflammatory cytokines.
			Nlrp3 and Casp2 are required for endoplasmic
141522	Casp2	26341399	reticulum stress-induced inflammation.
			The Mavs signalling pathway in non-myeloid cells is
007066		01454065	crucial for dsRNA-mediated natural killer cell
207066	Mavs	21454965	activation. Tyrosine phosphorylation of Mavs at amino acid
			residue Tyr9 is critical for the induction of Ifnb
207066	Mays	22844514	signalling. (Demonstrated in human)
			Upon infection with encephalitic Bunyavirus, RIG-
			I/MAVS signalling activates SARM1 to mediate
207066	Mavs	23499490	neuronal cell death.
			MAVS binds to VDAC1 to trigger viral-induced
207066	Mavs	23754752	apoptosis. Plasmodium RNA is a pathogen-associated molecular
			pattern (PAMP) capable of activating a type I IFN
			response via the cytosolic pattern recognition receptors
			Ifih1 and Mavs, as well as via transcription factors Irf3
207066	Mavs	24362933	and Irf7.
			Antiviral response to rotavirus in infected macrophages
207066	Mavs	26079065	is fully Mavs-dependent.
			Alveolar macrophages detect respiratory syncytial virus
207066	Mavs	25897172	(RSV) via the Mavs /Ddx58 pathway and are a major source of type I interferons upon RSV infection.
207000	1111115	23077172	Transgenic picornavirus RNA-dependent RNA
			polymerase (RdRP) expression in mice produces a
			quantitatively dramatic, sustained, effective antiviral
			interferon-stimulated genes (ISG) network, which
207066	Mavs	26633895	requires the MDA5-MAVS pathway.
207066	Mova	26246171	March5 modulates Mavs-mediated antiviral signalling,
207066	Mavs	202401/1	preventing excessive immune reactions. Baiap211 recruits Ube2i to sumoylate Pcbp2, which
			causes its cytoplasmic translocation during viral
			infection and the sumoylated Pcbp2 associates with
			Mavs to initiate its degradation, leading to
207066	Mavs	26348439	downregulation of antiviral responses.

			Pcbp2 synergizes with Pcbp1 in Mavs inhibition but Pcbp2 shows low basal expression with rapid induction after infection while Pcbp1 is stably and abundantly
188762	Pcbp2	22105485	expressed. (Demonstrated in human) Baiap211 recruits Ube2i to sumoylate Pcbp2, which causes its cytoplasmic translocation during viral infection and the sumoylated Pcbp2 associates with Mays to initiate its degradation, leading to
188762	Pcbp2	26348439	downregulation of antiviral responses. Baiap211 recruits Ube2i to sumoylate Pcbp2, which causes its cytoplasmic translocation during viral infection and the sumoylated Pcbp2 associates with
151396	Ube2i	26348439	Mavs to initiate its degradation, leading to downregulation of antiviral responses. Baiap211 recruits Ube2i to sumoylate Pcbp2, which causes its cytoplasmic translocation during viral infection and the sumoylated Pcbp2 associates with
208134	Baiap211	26348439	Mavs to initiate its degradation, leading to downregulation of antiviral responses. CASP1 activates NF-kappaB independent of its enzymatic activity and contributes to inflammation by
69618	CASP1	15039421	proteolysis of pro-IL1B (IL-1 beta) and RIPK2 activation of NF-kappaB and MAPK1. CASP1 is part of the inflammasome complex, along with pathogen-specific nucleotide oligomerization and binding domain (NOD)-like receptors (NLRs) and in some cases the scaffolding protein ASC. Formation of the membrane-associated inflammasome complex in
69618	CASP1	19124602	<ul> <li>murine macrophages, results in cleavage of cytosolic</li> <li>CASP1 substrates and cell death.</li> <li>CASP1 activity is required for discrimination between</li> <li>translocon-positive and -negative bacteria in bone-</li> <li>marrow derived cells and interleukin-1 receptor</li> <li>signalling. Activation of CASP1 by bacteria expressing</li> <li>Type 3 secretion systems allows for rapid recognition</li> <li>of bacteria expressing conserved functions associated</li> </ul>
69618	CASP1	20823203	with virulence. CASP1 clears intracellular bacteria in vivo independently of IL1B (IL-1-beta) and IL18 and establishes pyroptosis as an efficient mechanism of bacterial clearance by the innate immune system. CASP1-induced pyroptotic cell death releases bacteria from macrophages and exposes the bacteria to uptake
69618	CASP1	21057511	and killing by reactive oxygen species in neutrophils. CASP1 is a component of the inflammasome and is required for inflammation in acute pancreatitis.
69618	CASP1	21439959	(Demonstrated in murine model) (DASP1-dependent inflammatory cell death, or pyroptosis, is only induced by viable, but not heat- killed, E. coli. (Demonstrated in murine model)
69618	CASP1	21602824	,

(0(10		22022520	Naturally occurring variants of CASP1 differ considerably in structure and the ability to activate
69618	CASP1	22833538	IL1B. The precursor and mature forms of IL37 are secreted from activated cells upon inflammasome activation and
69618	CASP1	24481253	<ul><li>CASP1 processing of IL37 is important for its anti- inflammatory activity.</li><li>20-kDa IL1B generated from CASP1 cleaved pro-IL1B limits the available pro-IL1B for generation of CASP1</li></ul>
69618	CASP1	26324708	cleaved 17-kDa IL1B, thus reducing inflammation. IL1B is an important proinflammatory cytokine that activates monocytes, macropages, and neutrophils. IL1B processing during infection is a complex process in which the inflammasomes are only one of several
66519	IL1B	20195505	activation mechanisms. Mature IL1B production requires, in addition to the synthesis of pro-IL1B, cleavage of the precursor protein by the inflammatory CASP1 (Caspase-1) which
66519	IL1B	20401526	is controlled within the NLRP3 inflammasome. IL1B-producing conventional dendritic cells preserves and expands IL-22(+)AHR(+) immature human natural
66519	IL1B	20620944	killer cells in the secondary lymphoid tissue. IL1B acts as a growth factor for neutrophil progenitors and as a survival factor for mature neutrophils. In the absence of IKBKB, the IL1B production is enhanced and provides a compensatory mechanism for maintaining antibacterial defense when NFKB is inhibited. (Demonstrated in murine model)
66519	IL1B	21170027	minored. (Demonstrated in marine model)
			IL1B secretion in macrophages is regulated by autophagy by two mechanisms; sequestering of pro- IL1B in autophagosome during TLR stimulation, and processing, secretion of IL1B in a NLRP3- and TRIF- dependent manner.
66519	IL1B	21228274	-
			IL1B secretion is induced only during viable E. coli infection (as oppose to heat-killed E. coli or LPS); Viable bacteria specifically elicit cleavage of pro-IL1B. (Demonstrated in murine model)
66519	IL1B	21602824	
			IL1B derived from alveolar macrophages is the critical mediator which induces chemokine production in non- hematopoietic cells in the lung, resulting in swift and robust recruitment of infection-controlling neutrophils into the airways. (Demonstrated in murine model)
66519	IL1B	21270399	into the an ways. (Demonstrated in indime model)

IL1B secretion is tightly regulated by the redox status in myeloid cells. TLR engagement in monocytes induces ROS generation followed by a sustained antioxidant response and efficient IL1B secretion. In macrophages, the antioxidant systems are in an upregulated state, and therefore buffers the TLR induction of the redox response, which results in low IL1B processing and secretion.

IL1B is an important component of the cellular network involving macrophages and epithelial cells, which facilitates IL8 chemokine expression and aids neutrophil recruitment during pneumococcal pneumonia.

IL1B is an inflammatory cytokine that binds to its primary receptor, IL1R1, that then recruits the accessory protein IL1RAP to form a signallingcompetent heterotrimeric complex.

TLR8 plays a pathogenic role in disease whereby its expression is increased in patients with systemic arthritis and is correlated with the elevation of IL1B levels and disease status.

Protein-bound polysaccharide-K can activate the NLRP3 inflammasome and induce IL1B in a TLR2and NLRP3-dependent manner.

Interleukin-1 (IL1A/IL1B) plays a key role in the interaction between local vessel wall cells and invading cholesterol-triggered monocytes to multiply inflammation in the vessel wall. 25463072

IFNG interferes with the IL-1/NFKBIZ axis in  $\hat{I}^2$ glucan-activated dendritic cells and promotes T cellmediated immune responses with increased release of IFNG and IL22, and diminished production of IL17A.

CASP4 is a critical regulator of noncanonical inflammasome activation that initiates defence against bacterial pathogens in primary macrophages by mediating cell death and IL1A release

DEFB103A and RNASE7 are induced in human umbilical endothelial cells (HUVECs) by classical inflammatory cytokines such as: IFNG, IL1B and TNF. 25637949 Antibody-dependent enhancement (ADE) of Dengue virus serotype 2 (DENV-2) elevates mature IL1B secretion via SYK signalling pathway in primary 26032420 monocytes.

Differentiation of Type 3 innate lymphoid cells (ILC3) to IL7R(+) ILC1 is reversible whereas IL7R(+) ILC1 can differentiate to ILC3 in the presence of IL2, IL23A, and IL1B dependent on the transcription factor RORC, and this process is enhanced in the presence of retinoic 26187413 acid.

20-kDa IL1B generated from CASP1 cleaved pro-IL1B limits the available pro-IL1B for generation of CASP1 26324708 cleaved 17-kDa IL1B, thus reducing inflammation.

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66519 IL1B 22158745

66519 IL1B 22426547

66519 IL1B 24277153

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66519 IL1B 24323452

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IL1B 25474109

25964352

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TLR4 is activated by LPS and this recognition activates the Src family kinases, Src, Fyn and Yes, which in turn contribute to tyrosine phosphorylation of Zonula adherens proteins to open the endothelial paracellular pathway.

TLR4 binding to microbial ligands can be inhibited by CD180 and its helper molecule, LY86, via direct interactions with the TLR4 signalling complex.

TLR4 is involved in lipopolysaccharide (LPS) signaling and serves as a cell-surface co-receptor for CD14, leading to LPS-mediated NF-kappaB activation and subsequent cellular events.

TLR4-TLR6-Cd36 activation is a common molecular<br/>mechanism by which atherogenic lipids and amyloid-<br/>beta stimulate sterile inflammation.

TLR4 dimerize and enable rapid signal transduction<br/>against LPS stimulation on membrane-associated<br/>CD14-expressing cells.

TLR4 and TLR9 have both non-redundant and cooperative roles in lung innate responses during Gram-negative bacterial pneumonia and are both critical for IL-17 driven antibacterial host response.

TLR4 mediates LPS-induced muscle catabolism via coordinate activation of the ubiquitin-proteasome and the autophagy-lysosomal pathways. TLR4 activation by LPS induces C2C12 myotube atrophy via upregulating autophagosome formation and the expression of ubiquitin ligase atrogin-1/MAFbx and MuRE1

TLR4 transfection of eukaryotic host cells using bacterial vectors, or bactofection, was shown to reduce E. coli colonization in the kidney and the bladder in an animal model of urinary tract infection. (Demonstrated in murine model)

TLR4 is involved in the transmission of ER stress from tumour cells to macrophages, promoting a proinflammatory program in the tumour microenvironment, thus facilitating tumour progression. (Demonstrated in murine model)

TLR4 deficient murine macrophages results in the complete abrogation of TNF-alpha production during Leishmania panamensis infection. The endosomal TLR4 plays a crucial role in the activation of host macrophages and controlling the early stages of parasitic infection. (Demonstrated in murine model)

Epithelial TLR4 activation facilitates the transcytosis of non-cytolytic uropathogenic E. coli across intact collecting duct cell layers to invade the renal interstitium in experimental urinary tract infections.

82738TLR420826541MuRF1.<br/>TLR482738TLR420826541MuRF1.<br/>TLR482738TLR421442393in murine model)<br/>TLR4 is involved in<br/>tumour cells to minflammatory p<br/>microenvironment,<br/>Progression. (Demon<br/>TLR4 deficient mu

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TLR4:LY	Y96 functio	ons as int	racellula	ar LPS sens	or and
triggers	a unique	set of	LPS	responses	upon
recogniti	on of phag	ocytosed	bacteria	a in macrop	hages.
(Demonstrated in murine model)					

TLR4 on dendritic cell surfaces binds to HSPA14 and induces a robust Th1 response via the MAPK and NFkB signalling pathways. (Demonstrated in mouse)

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TLR4 recognizes Clostridium difficile surface layer proteins and induces the maturation of dendritic cells to activate the innate and adaptive immune response. (Demonstrated in mouse)

TLR4 and HSPD1 mediate myocardial ischemiaactivated innate immune signalling, which plays an important role in mediating apoptosis and inflammation during ischemia/reperfusion (I/R). (Demonstrated in murine model)

TLR4 and TLR2 are crucial for in vivo recognition of Chlamydia pneumoniae. Tlr4/2 double-deficient mice were unable to control pneumonia caused by C. 22096480 pneumoniae. (Demonstrated in mice)

TLR4 translocates to membrane lipid rafts in a ceramide-dependent manner in Helicobacter pylori 22354030 infected gastric epithelial cells.

TLR4 is involved in cell-cell contact signalling between activated apoptotic lymphocytes and dendritic 22396536 cells (DC) during the maturation of DCs.

Synthetic triacylated lipid A-molecules have the potent<br/>ability to selectively antagonize TLR4 and inhibit anti-<br/>bacterial immunity.

The poxviral protein A46 directly inhibits TLR4 signalling by disrupting receptor complex formation.

A human TLR4 polymorphism (D299G/T399I) impairs TLR4::LY96 dimerization and results in a dampened host response to bacterial lipids.

TLR4 is an important regulator of wound inflammation and is essential for early skin wound healing. (Demonstrated in mice)

TIR domain-contaning protein from Brucella melitensis, TcpB, disrupts the receptor-adaptor 24265315 interaction between TLR4 and TIRAP.

ECSIT binds to MAP3K7 and TRAF6 to form a complex that plays a pivotal role in activating TLR4mediated NF-kB signalling.

The TLR4/S100A8 axis is important in the activation of monocytes.

Endotoxin tolerance re-programs TLR4 signalling via suppression of PELI1, a positive regulator of MyD88and TIR domain-containing adapter inducing IFN-1<sup>2</sup> (TRIF)-dependent signalling that promotes K63-linked polyubiquitination of IRAK1, TBK1, and TAK1.

82738	TLR4	26610398	H. pylori infection induces the expression and activation of components of NLRP3 inflammasomes in neutrophils and this activation is independent of a functional type IV secretion system, TLR2 and TLR4. PELI3 is involved in endotoxin tolerance and functions as a negative regulator of TLR2/4 signalling.
82738	TLR4	26310831	TLR2 plays a critical role in the ability of innate immunity to determine M. pulmonis numbers in the lung, and early after respiratory infection TLR2
41789	TLR2	20505832	recognition of M. pulmonis triggers initial cytokine responses of host cells. TLR2 functions as a sensor of oxidation-associated molecular patterns, providing a key link connecting inflammation, oxidative stress, innate immunity and
41789	TLR2	20927103	angiogenesis. TLR1 :: TLR2 dimeric pairs recognize malarial glycosylphosphatidylinositols (GPI) to initiates
41789	TLR2	21439957	intracellular signalling and the production of pro- inflammatory cytokines. TLR2 recognizes Thermus aquaticus extracellular polysacchride, YT-1, and induces the production of
41789	TLR2	21454596	cytokines TNF and IL6 in peritoneal macrophages. (Demonstrated in murine model) TLR2::TLR6 synergistically interacts with TLR9 in lung epithelium to induce rapid pathogen killing, and
41789	TLR2	21482737	can be used as a therapeutic target to treat otherwise lethal pneumonia. TLR2 is activated by gut commensal microbe, Bacteroides fragilis, to establish host-microbial symbiosis by promoting immunological tolerance.
41789	TLR2	21512004	(Demonstrated in murine model) TLR2 and TNFSF11 signalling pathways are modulated by Porphromonas gingivalis to alter the differentiation states of osteoclasts resulting in bacteria-mediated bone loss. (Demonstrated in murine
41789	TLR2	21566133	model) TLR2 is expressed by Muller cells, principal glia of retina, and is responsible for generating robust bactericidal activity against Staphylococcus aureus and contributing to retinal innets dofined
41789	TLR2	21602496	contributing to retinal innate defence. TLR2 is required for rapid inflammasome activation in response to infection by cytosolic bacterial pathogens such as Francisella novicida. (Demonstrated in murine
41789	TLR2	21698237	model)

			TLR2-driven integration of inducible nitric oxide synthase (iNOS), Wnt-beta-Catenin and NOTCH1 signalling contributes to its capacity to regulate a battery of genes associated with T regulatory cell lineage commitment and towards modulation of defined set of effector functions in macrophages.
41789	TLR2	21862586	(Demonstrated in murine model) TLR2 directly recognizes glycogen, resulting in the activation of immunocytes such as macrophages to enhance the production of nitric oxide and
41789	TLR2	21873606	inflammatory cytokines. TLR2 and TLR4 are crucial for in vivo recognition of Chlamydia pneumoniae. Tlr2/4 double-deficient mice were unable to control pneumonia caused by C.
41789	TLR2	22096480	pneumoniae. (Demonstrated in mice) TLR2 signalling promotes protective vaccine- enhancing Th17 cell responses when cells are stimulated with early secreted antigenic target protein 6 (ESAT-6) expressed by the virulent Mycobacterium tuberculosis strain H37Rv but not by tuberculosis vaccine Bacillus Calmette-Gu à © rin (BCG).
41789	TLR2	22102818	(Demonstrated in mice) TLR2 recognizes Mycobacterium tuberculosis H37Rv cell surface lipoprotein MPT83, which induces the production of TNF, IL6, and IL12B cytokines by macrophages and upregulates macrophage function.
41789	TLR2	22174456	(Demonstrated in mouse) Mycobacterium abscessus glycopeptidolipid (GPL) prevents TLR2-mediate induction of IL8 and DEFB4A
41789	TLR2	22216191	in respiratory epithelial cells. Interaction of filamentous hemagglutinin (FHA) with TLR2 induces an innate immune response against Bordetella pertussis and the TLR2-binding domain of FHA may contribute to immunoprotection against
41789	TLR2	25353353	pertussis infection. Cutaneous bacteria can negatively regulate skin-driven immune responses by inducing Gr1(+)CD11b(+) myeloid-derived suppressor cells via TLR2-6
41789	TLR2	25456159	activation. Soluble TLR2 (sTLR2) generated by metalloproteinase activation inhibits TLR2-induced cytokine production
41789	TLR2	25531754	in THP-1 cell line. TLR10 is a functional receptor involved in the innate immune response to H. pylori infection and the TLR2/TLR10 heterodimer functions in H. pylori
41789	TLR2	25977263	lipopolysaccharide recognition. Human Cytomegalovirus (HCMV) miR-UL112-3p efficiently targets TLR2 during HCMV infection, resulting in the inhibition of TLR2-mediated NF ΰB
41789	TLR2	25955717	signalling.
41789 41789	TLR2 TLR2	26610398 26283364	H. pylori infection induces the expression and activation of components of NLRP3 inflammasomes in neutrophils and this activation is independent of a functional type IV secretion system, TLR2 and TLR4. Staphylococcal superantigen-like protein 3 (SSL3) interferes with TLR2 activation at two stages. First by binding to TLR2 and blocking ligand binding and second by interacting with an already formed TLR2- lipopeptide complex, thus preventing TLR heterodimerization and downstream signalling. PELI3 is involved in endotoxin tolerance and functions as a negative regulator of TLR2/4 signalling.
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41789	TLR2	26310831	PELI3 and other pellino isoforms are the E3 ubiquitin
59060 59060	PELI3 PELI3	17997719 25483963	ligases that mediate the IL-1-stimulated formation of K63-pUb-IRAK1 in cells, which may contribute to the activation of IKBKB and NF-kappaB, as well as other signalling pathways dependent on IRAK1 and IRAK4. Autophagy causes PELI3 degradation during Tlr4-signalling, subsequently inhibiting Il1b expression and impairing the hyperinflammatory phase during sepsis. PELI3 is involved in endotoxin tolerance and functions as a negative regulator of TLR2/4 signalling.
59060	PEL13	26310831	Signaling crosstalk during sequential Tlr4 and Tlr9 activation amplifies the inflammatory response of
156729	Tlr4	19923461	mouse macrophages.
156729	Tlr4	20385881	Tlr4 and Tlr2 activate murine macrophages and this activation is negatively regulated by a Lyn/PI3K module and promoted by SHIP1. Tlr4 transfection of eukaryotic host cells using bacterial
156729	Tlr4	21442393	vectors, or bactofection, was shown to reduce E. coli colonization in the kidney and the bladder in an animal model of urinary tract infection. Tlr4 is involved in the transmission of ER stress from tumour cells to macrophages, promoting a pro-
156729	Tlr4	21464300	inflammatory program in the tumour microenvironment, thus facilitating tumour progression. Tlr4 deficient murine macrophages results in the complete abrogation of TNF-alpha production during Leishmania panamensis infection. The endosomal Tlr4
156729	Tlr4	21518783	plays a cruical role in the activation of host macrophages and controlling the early stages of parasitic infection. Epithelial Tlr4 activation facilitates the transcytosis of non-cytolytic uropathogenic E. coli across intact collecting duct cell layers to invade the renal
156729	Tlr4	21615666	interstitium in experimental urinary tract infections.

			Tlr4:Ly96 functions as intracellular LPS sensor and
156729	Tlr4	21712422	triggers a unique set of LPS responses upon recognition of phagocytosed bacteria in macrophages. Tlr4 on dendritic cell surfaces binds to Hspa14 and
156729	Tlr4	21730052	induces a robust Th1 response via the MAPK and NFkB signalling pathways. Tlr4 recognizes Clostridium difficile surface layer
156729	Tlr4	21738466	proteins and induces the maturation of dendritic cells to activate the innate and adaptive immune response. Tlr4 and Hspd1 mediate myocardial ischemia-activated innate immune signalling, which plays an important
156729	Tlr4	21775438	role in mediating apoptosis and inflammation during ischemia/reperfusion (I/R). Tlr4 and Tlr2 are crucial for in vivo recognition of Chlamydia pneumoniae. Tlr4/2 double-deficient mice
156729	Tlr4	22096480	were unable to control pneumonia caused by C. pneumoniae. Tlr4 translocates to membrane lipid rafts in a ceramide-
156729	Tlr4	22354030	dependent manner in Helicobacter pylori infected gastric epithelial cells. (Demonstrated in human) Tlr4 is involved in cell-cell contact signalling between activated apoptotic lymphocytes and dendritic cells (DC) during the maturation of DCs. (Demonstrated in
156729	Tlr4	22396536	human) Synthetic triacylated lipid A-molecules have the potent
156729	Tlr4	22433865	ability to selectively antagonize Tlr4 and inhibit anti- bacterial immunity. (Demonstrated in human) The poxviral protein A46 directly inhibits Tlr4
156729	Tlr4	22593572	signalling by disrupting receptor complex formation. (Demonstrated in human) A human TLR4 polymorphism (D299G/T399I) impairs TLR4::LY96 dimerization and results in a dampened host response to bacterial lipids. (Demonstrated in
156729	Tlr4	22962435	human)
156729	Tlr4	22951730	Tlr4 is an important regulator of wound inflammation and is essential for early skin wound healing. Milk oligosaccharide sialyl( $\hat{1} \pm 2,3$ )lactose modulates
156729	Tlr4	24101501	mucosal immunity by inducing inflammation through TLR4 signaling Itgam (Cd11b) fine tunes the balance between adaptive and innate immune responses initiated by LPS by modulating the trafficking and signalling functions of
156729	Tlr4	24423728	Tlr4 in a cell-type-specific manner. Nod1 and Nod2 synergize with Tlr4 in dendritic cells to increase IL12 production and enhance invariant natural killer T (iNKT) cell activation, and are important regulators of the IFN gamma response by iNKT cells during S. typhimurium and L.
156729	Tlr4	24163408	monocytogenes infections. Rab8a interacts with Pik3cg to regulate Akt signalling
156729	Tlr4	25022365	generated by surface Tlr4.

		High-potency Tlr4 agonists can act as clinically useful vaccine adjuvants by selectively activating Ticam1- dependent immunostimulatory signalling events and only weakly activating potentially harmful Myd88-
Tlr4	25389373	dependent inflammatory responses.
Tlr4	25448706	Myd88 and Ticam1 pathways differently regulate Tlr4- induced immune responses in B cells. Autophagy causes PELI3 degradation during Tlr4- signalling, subsequently inhibiting Il1b expression and
Tlr4	25483963	impairing the hyperinflammatory phase during sepsis. Tmem126a upregulates genes involved in antigen
Tlr4	25549946	presentation; such as Icam1, MHC II, Cd86 and Cd40, via the Tlr4 signal transduction pathway. Ticam1 but not Myd88 signalling is critical for the Trl4 protective adjuvant effect in neonates; where Ticam1(- /-) but not Myd88(-/-) neonates are highly susceptible
Tlr4	25548220	to Escherichia coli peritonitis and bacteremia. Wdfy1 is a crucial adaptor protein in the Tlr3/4 signalling pathway. Wdfy1 interacts with Tlr3 and Tlr4 and mediates the recruitment of Ticam1 to these
Tlr4	25736436	receptors. Lipopolysaccharide-mediated myeloid Anpep (CD13) expression governs internalization of Tlr4 and negatively regulates Tlr4 signalling, thereby balancing the innate response by maintaining the inflammatory
Tlr4	25801433	equilibrium critical to innate immune regulation. Psen2 deficiency is paralleled by reduced transcription of Tlr4 mRNA and loss of LPS-induced Tlr4 mRNA
Tlr4	26081153	transcription regulation. Peli3 is involved in endotoxin tolerance and functions as a negative regulator of Tlr2/4 signalling.
Tlr4	26310831	Peli3 is involved in endotoxin tolerance and functions as a negative regulator of Tlr2/4 signalling.
Peli3	26310831	Atf7, a stress-response transcription factor, mediates epigenetic changes in macrophages involved in innate immunological memory in response to

			immunological memory in response to
189070	Atf7	26322480	lipopolysaccharide.
			MIR223 negatively regulates NLRP3 inflammasome
127311	<b>MIR223</b>	22984082	activity.
			MIR223 regulates the NLRP3 inflammasome and IL-
127311	<b>MIR223</b>	22984081	1Î <sup>2</sup> production.
			MIR223 regulates macrophage function by modulating
			cytokine production and NF-ΰB activation through
			inhibition of RELA phosphorylation and nuclear
127311	MIR223	26296289	translocation.
			Soluble COLEC12 can recognize Aspergillus
			fumigatus leading to activation of the alternative
285	COLEC12	26290605	pathway of complement.

47294	TLR3	19074283	TLR3-type II IFN signalling cooperates with the RIG- I/MDA5-type I IFN axes for efficient innate antiviral immune responses.
47294	TERS	19074283	TLR3-dependent antiviral pathway is negatively regulated by activated F2RL1 leading to blunted expression of TLR3/IRF3 driven genes, as well as
47294	TLR3	19865078	activation of IRF3 and STAT1. TLR3, TLR2, or TLR4 cooperate with proteinase- activated receptors (PARs) for activation of nuclear
47294	TLR3	19865078	factor-kappaB-dependent signalling in mucosal epithelial cell lines. TLR3-induced proapoptotic signalling involves
47294	TLR3	20019748	TICAM1 (TRIF)-dependent activation of CASP8 and is under the control of inhibitor of apoptosis proteins (IAPs) in melanoma cells. TLR3 is activated by poly(I:C) and this induces
47294	TLR3	11607032	cytokine production through a signalling pathway dependent on MyD88. TLR3-mediated activation of NF-kappaB and IRF3
47294	TLR3	14982987	diverges at Toll-IL-1 receptor domain-containing adapter 1 (TICAM1) inducing IFN-beta. TLR3 dimerizes when it binds dsRNA and this is essential for ligand binding. Although the three TLR3 contact sites individually interact weakly with their
47294	TLR3	20861016	<ul><li>binding partners, together they form a high affinity ligand-receptor complex.</li><li>TLR3 deletion dramatically enhanced the development of elastic lamina damage after collar-induced injury,</li></ul>
47294	TLR3	21220319	indicating that TLR3 signalling plays a protective role in arterial vessel wall. TLR3 activation by Poly(I:C) in the endothelial cells induces Poly(I:C) dose- and time-dependent cell apoptosis. Specifically, TLR3 stimulation triggered the
47294	TLR3	21367858	signalling of both extrinsic and intrinsic apoptotic pathways. TLR3 requires proteolytic processing in endolysosome by asparagine endopeptidase and cathepsin in the
47294	TLR3	21402738	endolysosome to initiate signalling in response to DNA. (Demonstrated in murine model) TLR3 expression is inducible by LPS via TLR4-
47294	TLR3	21498625	MYD88-IRAK-TRAF6-NFKB dependent signalling pathway. TLR3 is necessary to establish an antiviral state in hepatocytes infected with hepatitis C Virus. HCV
47294	TLR3	21695051	envelope proteins counteract the antiviral host defence by inhibiting the expression of TLR3.
47294	TLR3	22016778	TLR3 signalling is enhanced by the presence of viral double-strand RNA-binding proteins. TLR3-TICAM1-mediated signalling pathway plays an
47294	TLR3	22072781	essential role in the anti-viral response against poliovirus infection. (Demonstrated in mice)

			TLR3 is constitutively expressed in spermatogonia and spermatocytes, and has the ability to activate anti-viral
47294	TLR3	22262694	responses. (Demonstrated in mice) Upon engagement with its ligand, dsRNA, TLR3
47294	TLR3	22421964	possesses the ability to recruit CASP8 and RIPK1 to induce apoptosis. Activation of TLR3 with poly(I:C) mediates antiviral
47294	TLR3	22754655	immunity that diminishes coronavirus production in macrophages. (Demonstrated in mice) Upregulation of TLR3 in intestinal epithelia during infancy may contribute to age-dependent susceptibility
47294	TLR3	22570612	to rotavirus infection. (Demonstrated in mice) TLR3 activation differentially regulates phagocytosis
47294	TLR3	22986631	of bacteria and apoptotic neutrophils by peritoneal macrophages. (Demonstrated in mice) TLR3 is more efficiently activated by high molecular
47294	TLR3	23035017	mass than by low molecular weight poly(I:C). TLR3-mediated antibody response to Chikungunya virus plays a key role in its infection, replication and
47294	TLR3	25452586	pathology. Intracellular/endocytic TLR3 interacts with SCARF 1 in the presence of Poly I:C to boost TLR3-mediated
47294	TLR3	25641411	inflammatory signalling and stimulate cytokine production in macrophages. Signalling through both DDX58 and TLR3 is
47294	TLR3	25880109	important for interferon induction by influenza A virus in alveolar epithelial cells. Bluetongue virus activates TLR3/interferons signalling
47294	TLR3	26296370	<ul><li>pathway resulting in the inhibition of human immunodeficiency virus in macrophages.</li><li>CLEC4E, a C-type lectin receptor, is a pattern recognition receptor critical for immune responses to fungi. CLEC4E is coupled to SYK kinase and signals via CARD9 to activate NFKB, which in turns induces both innate and adaptive immunity.</li></ul>
17212	CLEC4E	21267996	
17212	CLEC4E	24733387	Glycerol monomycolate is a unique ligand for CLEC4E (Mincle). Cholesterol crystals are an endogenous ligand for
17212	CLEC4E	26296894	CLEC4E and their binding activates innate immune responses. Traf6 is a E3 ubiquitin ligase that activates NFKB pathway in response to innate and adaptive immunity stimuli. Traf6 protein contains a highly conserved TRAF-C domain that contributes to oligomerization and its interaction to upstream signalling molecules, and a RING domain dimerization interface that is functionally important for ubiquitination and the activation of NFKB.
193222	Traf6	21185369	

Traf6 is polyubiquitinated and disassembled during endotoxin tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of antiinflammatory or anti-microbial mediators.

193222	Traf6	21220427	initialinitatory of anti-inicrobial mediators.
193222	Traf6	22033459	Traf6 is degraded in the proteasome upon TLR stimulation in macrophages. The binding of MAVS to Traf2, Traf5, and Traf6 is dependent on virus infection and MAVS
193222	Traf6	23951545	<ul> <li>polymerization. The TRAF proteins promote ubiquitination that recruits IKBKG binding to the MAVS signalling complex.</li> <li>Mir146 attenuates sepsis-induced cardiac dysfunction by preventing NF Î °B activation, inflammatory cell infiltration, and inflammatory cytokine production via</li> </ul>
193222	Traf6	26048146	targeting of Irak1 and Traf6 in both cardiomyocytes and inflammatory monocytic cells Intracellular Sef/IL-17R (SEFIR) domain of Il17rd targets TIR adaptor proteins Myd88, Tirap, Ticam1,
193222	Traf6	25808990	Ticam2 and Traf6 to inhibit TLR downstream signalling. Trim12c interacts with Traf6, a key protein in pathogen recognition receptor signalling, and reciprocally
193222	Traf6	26503954	enhances its ubiquitination, leading to cooperative activation of IFN and NF-kB pathways. Usp25 physically associates with Traf3 and Traf6 after infection by RNA or DNA viruses and promotes innate antiviral responses by protecting virus-induced
193222	Traf6	26305951	<ul><li>proteasome-dependent or independent degradation of Traf3 and Traf6.</li><li>Traf3 is a component of Toll/interleukin-1 receptor (TIR) signalling complexes that is recruited along with Traf6. Traf3 is essential for the induction of type I</li></ul>
172558	Traf3	16306937	interferons (IFN) and the anti-inflammatory cytokine interleukin 10, but is dispensable for expression of pro- inflammatory cytokines. Traf3 is a highly versatile regulator that positively controls type I interferon production, and negatively
172558	Traf3	21660053	regulates MAP kinase activation and alternative NFkB signalling. Upon sensing dsRNA or dsDNA, Traf3 interacts with ER-to-Golgi transport proteins to induce Mavs-
172558	Traf3	22792062	associated innate immune responses. (Demonstrated in human) Traf3 expressed in myeloid cells regulates immune
172558	Traf3	25422508	responses in myeloid cells and acts to inhibit inflammation and tumor development.

172558	Traf3	26305951	Usp25 physically associates with Traf3 and Traf6 after infection by RNA or DNA viruses and promotes innate antiviral responses by protecting virus-induced proteasome-dependent or independent degradation of Traf3 and Traf6. Usp25 physically associates with Traf3 and Traf6 after infection by RNA or DNA viruses and promotes innete
171582	Usp25	26305951	<ul> <li>infection by RNA or DNA viruses and promotes innate antiviral responses by protecting virus-induced proteasome-dependent or independent degradation of Traf3 and Traf6.</li> <li>Irf1 transcriptionally inhibits the Il23a through the ISRE element and reduce the severity of chronic intestinal inflammation caused by LPS.</li> </ul>
172251	Irf1	21097874	
172251	Irfl	22266972	Irfl promotes immune cell apoptosis and inhibits autophagy in a murine endotoxemia model. Irfl is an essential regulator of the host innate antiviral response in the brain by limiting viral replication at
172251	Irf1	24675692	later stages of infection, but is not involved in the rapid induction of IFN. Nlrp3 deficiency protects mice from the development of type 1 diabetes by suppressing Th1 responses and
172251	Irfl	26305961	impairing T-cell migration to pancreatic islets through the down-regulation of chemokine expression (Ccl5, Cxcl10, Irf1) in islets. Cxcl10 exert direct antimicrobial effects in vitro against Bacillus anthracis spore and bacilli in a receptor-independent manner and contributes to pulmonary innate immunity.
179916	Cxcl10	21124994	Cxcl10 concentration in blood increases during neonatal polymicrobial sepsis, and the blockade of Cxcl10 not only worsens recruitment and phagocytic function of macrophages, but also the survival of neonatal mice.
179916	Cxcl10	21518789	Nlrp3 deficiency protects mice from the development of type 1 diabetes by suppressing Th1 responses and impairing T-cell migration to pancreatic islets through
179916	Cxcl10	26305961	the down-regulation of chemokine expression (Ccl5, Cxcl10, Irf1) in islets. Nlrp3 deficiency protects mice from the development of type 1 diabetes by suppressing Th1 responses and impairing T-cell migration to pancreatic islets through
205509	Ccl5	26305961	the down-regulation of chemokine expression (Ccl5, Cxcl10, Irf1) in islets. ITGB1 along with ITGA3 is a novel regulator for the recognition of bacterial lipopeptides. ITGB1/ITGA3 integrin regulates endosomal Toll-like receptor (TLR)-
68302	ITGB1	20877569	2/TLR1 signalling, serving as a mechanism for modulating inflammatory responses.

68302	ITGB1	26288256	IFNG primes mast cells for enhanced anti-bacterial and pro-inflammatory responses to Staphylococcus aureus, partially mediated by ITGB1.
			The combined treatment of IFNG with 1,25- dihydroxyvitamin D3 (1,25-D3) synergistically enhances nitric oxide (NO) synthesis and NOS2 expression induced by Mycobacterium tuberculosis
45916	IFNG	20157607	<ul><li>(MTB) or by its purified protein derivatives in human monocyte-derived macrophages.</li><li>IFNG mediates DUOX2 dual oxidase expression via a STAT-independent signalling pathway and providing insights into a novel IFNG signalling pathway with</li></ul>
45916	IFNG	20381453	potential importance for regulation of host defence responses. IFNG is crucial for immunity against intracellular pathogens and for tumour control and it is produced
45916	IFNG	17981204	predominantly by natural killer (NK) and natural killer T (NKT) cells as part of the innate immune response. IFN gamma creates a primed chromatin environment in macrophages to augment TLR-induced gene
45916	IFNG	24012417	transcription. IFNG interferes with the IL-1/NFKBIZ axis in $\hat{1}^2$ -glucan-activated dendritic cells and promotes T cell-
45916	IFNG	25474109	mediated immune responses with increased release of IFNG and IL22, and diminished production of IL17A. Primary $\hat{1}^{3}\hat{1}'$ T cells provide an early source of IFNG during dengue virus (DV) infection and target DV-infected cells. Monocytes also participate as accessory
45916	IFNG	25732728	cells that sense DV infection and amplify the cellular immune response in an IL18-dependent manner. DEFB103A and RNASE7 are induced in human umbilized and the light cells (ILIN/ECs) by classical
45916	IFNG	25637949	umbilical endothelial cells (HUVECs) by classical inflammatory cytokines such as: IFNG, IL1B and TNF. IFNG primes mast cells for enhanced anti-bacterial and pro-inflammatory responses to Staphylococcus aureus,
45916	IFNG	26288256	partially mediated by ITGB1. DDX60L is an important effector protein of the innate
43931	DDX60L	26269178	immune response against hepatitis C virus. Histone deacetylase 2 (HDAC2) is part of a repressor complex, along with key components that include HDAC1, RE-1 silencing transcription factor (REST), co-repressor of REST (CoREST), and lysine-specific demethylase (LSD) 1. The HDAC/CoREST/REST/LSD1 repressor complex is a
95591	HDAC2	20798038	significant component of host innate immunity. TET2 selectively mediates active repression of IL6 transcription via NFKBIZ and HDAC2 during
95591	HDAC2	26287468	inflammation resolution in innate myeloid cells, including dendritic cells and macrophages.
48252	NFKBIZ	16513645	Inhibits the DNA binding of RELA and NFKB1

			A key regulator of IL-6 production in human monocytes and plays an important role in both TLR
48252	NFKBIZ	19783680	and NOD-like receptor ligand induced inflammation IFNG interferes with the IL-1/NFKBIZ axis in $\hat{1}^2$ -glucan-activated dendritic cells and promotes T cell-
48252	NFKBIZ	25474109	mediated immune responses with increased release of IFNG and IL22, and diminished production of IL17A. TET2 selectively mediates active repression of IL6 transcription via NFKBIZ and HDAC2 during
48252	NFKBIZ	26287468	<ul><li>inflammation resolution in innate myeloid cells, including dendritic cells and macrophages.</li><li>IL6 trans-signalling via STAT3 is a critical modulator of LPS-driven pro-inflammatory responses through cross-talk regulation of the TLR4/Mal signalling pathway broader mechanism that regulates the severity of the host inflammatory response.</li></ul>
9462	IL6	21148800	IL6 synthesis is regulated by the opposing effects of prostaglandin (PG)E(2) and PGD(2) in human chondrocytes. IL6 synthesis is increased by PGE2 and decreased by PGD2 through the modulation of TLR4
9462	IL6	22096605	synthesis. Activation of either TLR4 or TLR2/6 significantly increased IL6 expression by U937 mononuclear cells. Co-activation of TLR4 and TLR2/6, led to a further
9462	IL6	22030478	augmentation on IL-6 expression. IL6 is strategically upregulated by virulent Mycobacterium tuberculosis to inhibit the induction of
9462	IL6	22426116	innate immunity.
9462	IL6	23735697	Hyperglycemia abrogates the ability of IL6 to induce neutrophil extracellular traps. IFN gamma creates a primed chromatin environment in
9462	IL6	24012417	macrophages to augment TLR-induced IL6 transcription TET2 selectively mediates active repression of IL6 transcription via NFKBIZ and HDAC2 during
9462	IL6	26287468	inflammation resolution in innate myeloid cells, including dendritic cells and macrophages. TET2 selectively mediates active repression of IL6 transcription via NFKBIZ and HDAC2 during
32690	TET2	26287468	inflammation resolution in innate myeloid cells, including dendritic cells and macrophages. Tet2 selectively mediates active repression of Il6 transcription via Nfkbiz and Hdac2 during
143980	Hdac2	26287468	inflammation resolution in innate myeloid cells, including dendritic cells and macrophages. Tet2 selectively mediates active repression of Il6 transcription via Nfkbiz and Hdac2 during
165943	Nfkbiz	26287468	inflammation resolution in innate myeloid cells, including dendritic cells and macrophages.

			Il6 trans-signaling via Stat3 is a critical modulator of LPS-driven pro-inflammatory responses through cross-talk regulation of the Tlr4/Mal signaling pathway as a broader mechanism that regulates the severity of the host inflammatory response.
144970	I16	21148800	
144970	Il6	22096605	Il6 synthesis is regulated by the opposing effects of prostaglandin (PG)E(2) and PGD(2) in human chondrocytes. Il6 synthesis is increased by PGE2 and decreased by PGD2 through the modulation of Tlr4 synthesis. (Demonstrated in human) Activation of either Tlr4 or Tlr2/6 significantly increased Il6 expression by U937 mononuclear cells. Co-activation of Tlr4 and Tlr2/6, led to a further
144970	Il6	22030478	augmentation on Il6 expression. (Demonstrated in human)
			Il6 is strategically upregulated by virulent Mycobacterium tuberculosis to inhibit the induction of
144970	I16	22426116	innate immunity. (Demonstrated in human)
144970	Il6	23359591	Mycobacterium tuberculosis regulates host IL6 production to inhibit type I interferon-signalling.
144970	по	23337371	Il6 causes compromised tissue repair by shifting acute inflammation into a more chronic profibrotic state
144970	I16	24412616	through induction of T helper type1 cell responses as a consequence of recurrent inflammation. Ybx1 controls intracellular II6 mRNA levels in a cell
			type-specific manner, leading to functions that are
1 4 4 9 7 9			dependent on the extracellular and intracellular
144970	Il6	25398005	distribution of Ybx1. Cfp plays a role in intestinal homeostasis in response to
			an infectious challenge to activate Hc (C5a), which in
144070	ЦС	0.570.510.5	turn provides protection through Il6 expression by the
144970	Il6	25725105	epithelium. Defb1 is important for the control of early mucosal
			Candida infection and plays a critical role in the
144070	116	25595775	induction of innate inflammatory mediators including,
144970	Il6	25595775	Il1b, Il6, Cxcl1, Il17a, and Il17f. Tet2 selectively mediates active repression of Il6
			transcription via Nfkbiz and Hdac2 during
144970	I16	26287468	inflammation resolution in innate myeloid cells, including dendritic cells and macrophages.
144770	по	20207400	Tet2 selectively mediates active repression of Il6
			transcription via Nfkbiz and Hdac2 during
195027	Tet2	26287468	inflammation resolution in innate myeloid cells, including dendritic cells and macrophages.
198027	1002	2020,100	March5 is a mitochondrial ubiquitin ligase that
161269	M. 17	21/25525	positively regulates Tlr7 signalling. March5 interacts
161368	March5	21625535	with Tank to induce NFkB-mediated gene expression. March5 modulates Mavs-mediated antiviral signalling,
161368	March5	26246171	preventing excessive immune reactions.

			MARCH5 is a mitochondrial ubiquitin ligase that positively regulates TLR7 signalling. MARCH5
82450	MARCH5	21625535	interacts with TANK to induce NFkB-mediated gene expression.
82450	MARCH5	26246171	MARCH5 modulates MAVS-mediated antiviral signalling, preventing excessive immune reactions. Mitophagy-inducing proteins Bnip3 and Bnip31 play a regulatory role in the generation of robust natural killer
175760	Bnip31	26253785	cell memory. Mitophagy-inducing proteins Bnip3 and Bnip31 play a
269444	Bnip3	26253785	regulatory role in the generation of robust natural killer cell memory. Isg15 expression is induced during a hepatitis C virus infection in an Irf3-dependent manner. Isg15 acts as a negative modulator of the RIG-I pathway by mediating the level of Ddx58 (RIG-I) ubiquitination.
137564	Isg15	22022264	(Demonstrated in human)
137564	Isg15	26259872	Isg15 protects against Listeria monocytogenes infection. ISG15 (ISGylation) post-translational modification is mediated by a sequential reaction similar to
84312	ISG15	19043203	ubiquitination and has been shown to negatively regulate the NF-kappaB pathway. ISG15 expression is induced during a hepatitis C virus infection in an IRF3-dependent manner. ISG15 acts as
84312	ISG15	22022264	a negative modulator of the RIG-I pathway by mediating the level of DDX58 (RIG-I) ubiquitination. ISG15 does not directly alter human rhinovirus replication but modulates immune signalling via the viral sensor protein DDX58 to impact production of CXCL10, which has been linked to innate immunity to
84312	ISG15	24448099	viruses. Human rhinovirus infection of epithelial cells induces the expression and secretion of ISG15, which
84312	ISG15	24448099	modulates immune responses via effects on DDX58, and by regulating CXCL10 production.
84312	ISG15	26259872	ISG15 protects against Listeria monocytogenes infection. During sepsis-induced generation of myeloid-derived
159848	Cdkn1a	26259914	suppressor cells, transcription factor Nfia represses Cdkn1a to arrest differentiation of Gr1+ CD11b+ cells. During sepsis-induced generation of myeloid-derived
164923	Nfia	26259914	suppressor cells, transcription factor Nfia represses Cdkn1a to arrest differentiation of Gr1+ CD11b+ cells. IFNB1 has an essential role in the anti-viral response
53485	IFNB1	20174559	and optineurin (OPTN) has a role in the inhibition of virus-triggered IFNB1 induction. Viral RNA-induced IFNB1 production is suppressed
53485	IFNB1	20501842	by oncogenic RAS through negative regulation of RIG- I signalling, leading to promotion of virus spread.

			IFNB1 expression is inhibited by influenza A virus polymerase by binding to IFN-beta promoter stimulator
53485	IFNB1	20699220	1 (MAVS). IFNB1 is a type I interferon (IFN-I) and the IFN-I family comprises a wide number of cytokines with
53485	IFNB1	20298124	different modulatory effects on angiogenesis, cell growth, fibrosis, apoptosis and autoimmunity. IFNB1 deficiency results in a partial suppression of the sterol pathway in macrophages during viral infections, thereby linking the regulation of lipid metabolism
53485	IFNB1	21408089	pathway with interferon anti-viral defence responses. (Demonstrated in murine model) IFNB1 secretion is greater upon viable E. coli infection in comparison to heat killed E. coli vaccine or LPS. The induction of IFNB1 is dependent on TICAM1- IRF3 signalling. (Demonstrated in murine model).
53485	IFNB1	21602824	
			IFNB1 expression pattern during viral infection is a highly stochastic process influenced by cell-to-cell variability in viral induction processes. (Demonstrated
53485	IFNB1	22291574	in mice) IFNB1 production is fundamental to the efficient control of Listeria monocytogenes during the early innate phase of infection. NK cells treated with IFNB1 during early infection were able to reduce bacterial titer
53485	IFNB1	22912878	in the spleen and significantly improve survival of infected mice. (Demonstrated in mouse) Macrocyclic NS3-4A resistance-associated amino acid variants (RAVs) with substitutions at residue D168 of the hepatitis C virus protease result in an increased capacity of NS3-4A to cleave MAVS and suppress
53485	IFNB1	25463536	IFNB1 induction. Coronavirus engages papain-like proteases to escape
53485	IFNB1	25505178	from the innate antiviral response of the host by inhibiting TP53-IRF7-IFNB1 signalling. Hepatitis B virus (HBV) polymerase inhibits
53485	IFNB1	25505063	TMEM173-stimulated IRF3 activation and IFNB1 induction. ELAVL1 is required for the stabilization of IFNB1 mRNA, and suppression of ELAVL1 leads to impaired
53485	IFNB1	25678110	expression of IFNB1 in response to poly(I:C) treatment. IFNB1 selectively restricts the transcriptional responses mediated by both the TLRs and the NOD-
53485	IFNB1	26202980	like receptors in Salmonella enterica serovar Typhimurium infection in macrophages. Reversible arginine methylation of TRAF6 is regulated
70010	JMJD6	26221041	by PRMT1 and JMJD6 and this in turn regulates TRAF6-dependent TLR signalling.

PRMT1 is a protein arginine methyltransferase and a novel and crucial negative regulator of STAT1 activation that controls PIAS1-mediated repression by arginine methylation.

63299	PRMT1	19136629	
63299	PRMT1	26221041	Reversible arginine methylation of TRAF6 is regulated by PRMT1 and JMJD6 and this in turn regulates TRAF6-dependent TLR signalling.
05277		20221041	Differentiation of Type 3 innate lymphoid cells (ILC3) to IL7R(+) ILC1 is reversible whereas IL7R(+) ILC1 can differentiate to ILC3 in the presence of IL2, IL23A, and IL1B dependent on the transcription factor RORC, and this process is enhanced in the presence of retinoic
102434	RORC	26187413	acid. IL23A expression is regulated by MAP3K8 in lipopolysaccharide (LPS)-stimulated macrophages
40891	IL23A	20405269	through ERK activation. IL23A suppresses innate immune response independently of IL17A during carcinogenesis and
40891	IL23A	20404142	metastasis (shown in mice). IL23A is a LPS induced gene, and its expression in
			macrophages correlate with the severity of chronic intestinal inflammation. IL23A is transcriptionally inhibited by the binding of IRF1 to the ISRE element and serve as a homeostatic checkpoint in chronic
40891	IL23A	21097874	intestinal inflammation (shown in mice). Differentiation of Type 3 innate lymphoid cells (ILC3) to IL7R(+) ILC1 is reversible whereas IL7R(+) ILC1 can differentiate to ILC3 in the presence of IL2, IL23A, and IL1B dependent on the transcription factor RORC, and this process is enhanced in the presence of retinoic
40891	IL23A	26187413	acid. Differentiation of Type 3 innate lymphoid cells (ILC3) to IL7R(+) ILC1 is reversible whereas IL7R(+) ILC1 can differentiate to ILC3 in the presence of IL2, IL23A, and IL1B dependent on the transcription factor RORC, and this process is enhanced in the presence of retinoic
36892	IL2	26187413	acid. Differentiation of Type 3 innate lymphoid cells (ILC3) to IL7R(+) ILC1 is reversible whereas IL7R(+) ILC1 can differentiate to ILC3 in the presence of IL2, IL23A, and IL1B dependent on the transcription factor RORC, and this process is enhanced in the presence of retinoic
16161	IL7R	26187413	acid. Macrophage iron regulatory proteins Aco1 and Ireb2 are protective against Salmonella by promoting the induction of Lcn2, a host antimicrobial factor that inhibits bacterial uptake of iron-laden siderophores, and
160059	Len2	26190773	by suppressing the ferritin iron pool.

			Macrophage iron regulatory proteins Aco1 and Ireb2 are protective against Salmonella by promoting the induction of Lcn2, a host antimicrobial factor that inhibits bacterial uptake of iron-laden siderophores, and
167179	Ireb2	26190773	by suppressing the ferritin iron pool. Macrophage iron regulatory proteins Aco1 and Ireb2 are protective against Salmonella by promoting the induction of Lcn2, a host antimicrobial factor that inhibits bacterial uptake of iron-laden siderophores, and
135868	Acol	26190773	by suppressing the ferritin iron pool. TMEM173 (STING) is an endoplasmic reticulum (ER) receptor that facilitates interferon (IFN) induction by binding to DDX58 (RIG-I) and to subunits of TRAP complex that facilitates translocation of proteins into
47893	TMEM173	18724357	the ER following translation. TMEM173 is a critical mediator of virus-triggered type I IFN signalling and a critical mediator of virus-
47893	TMEM173	18818105	triggered IRF3 activation. TMEM173 is essential for host defence against DNA
47893	TMEM173	19776740	<ul><li>pathogens such as HSV-1 and facilitates the adjuvant activity of DNA-based vaccines.</li><li>TMEM173 is an adaptor protein that links virus-sensing receptors to IRF3 activation. RNF5 negatively regulates this virus-triggered signaling by targeting</li></ul>
47893	TMEM173	19285439	<ul><li>TMEM173 for ubiquitination and degradation at the mitochondria.</li><li>TMEM173 is required for double stranded DNA-triggered innate immune responses where, upon sensing dsDNA, TMEM173 moves from the endoplasmic reticulum (ER) to the Golgi apparatus and</li></ul>
47893	TMEM173	19926846	finally reaches the cytoplasmic punctate structures to assemble with TANK-binding kinase 1 (TBK1). TMEM173 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA
47893	TMEM173	21820332	and in the subsequent induction of type I IFNs. (Demonstrated in mouse) TMEM173 activates STAT6 during viral infection to
47893	TMEM173	22000020	<ul><li>induce genes responsible for immune cell homing.</li><li>(Demonstrated in mice)</li><li>TMEM173 is cleaved by dengue viral protease to</li></ul>
47893	TMEM173	22761576	suppress IRF3 activation and subvert antiviral immunity. Dengue viral NS2B3 protease complex selectively
47893	TMEM173	23055924	targets TMEM173 (STING) for degradation to inhibit type I IFN production in human dendritic cells. TMEM173 (STING) is targeted by hepatitis C viral
47893	TMEM173	23542348	protease to disrupt interferon signalling. Cyclic-di-GMP-induced levels of IFI16 suppress the
47893	TMEM173	24131791	expression of TMEM173 (STING).

			In herpes simplex virus 1 (HSV-1) infected cells, the stability and function of IFI16 and TMEM173 are dependent on cell derivation and the functional integrity of HSV 1 proteins ICP0 and HS2 proteins
47893	TMEM173	24449861	integrity of HSV-1 proteins ICP0 and US3 protein kinase. After viral infection, ELF4 binds to TMEM173 (STING) and induces type I interferon. ELF4 is critical
47893	TMEM173	24185615	for host antiviral defense. The end result of the interplay between TMEM173 (STING), IFI16, and herpes simplex virus 1 (HSV-1) is determined by the genotype of the infected cells and the functional integrity of HSV-1 proteins infected cell
47893	TMEM173	24449861	protein 0 (ICP0) and US3 protein kinase. Familial TMEM173 mutation is associated with inflammatory lupus-like manifestations.
47893	TMEM173	25401470	Cytosolic RNA:DNA hybrids are sensed by the
47893	TMEM173	25425575	MB21D1-TMEM173 (cGAS-STING) pathway of the innate immune system. Hepatitis B virus (HBV) polymerase inhibits TMEM173-stimulated IRF3 activation and IFNB1
47893	TMEM173	25505063	induction. Upon cytoplasmic DNA stimulation, the endoplasmic
47893	TMEM173	25526307	reticulum protein AMFR is recruited to and interacts with TMEM173 in an INSIG1-dependent manner. Stimulation of TMEM173-dependent IRF3 activation by ultraviolet radiation is due to apoptotic signalling-
47893	TMEM173	25792739	dependent disruption of ULK1, a pro-autophagic protein that negatively regulates TMEM173. 4-(2-chloro-6-fluorobenzyl)-N-(furan-2-ylmethyl)-3- oxo-3,4-dihydro-2H-benzo[b][1,4]thiazine-6- carboxamide (G10) requires STING to trigger IRF3/IFN-associated transcription in human fibroblasts and subsequently blocking replication of Chikungunya
47893	TMEM173	26646986	virus, Venezuelan Encephalitis virus, and Sindbis virus. Viral interferon regulatory factor 1 (vIRF1), targets TMEM173 by preventing it from interacting with TBK1, thereby inhibiting TMEM173's phosphorylation and concomitant activation, resulting in an inhibition
47893	TMEM173	26199418	of the DNA sensing pathway. TBK1 alternative splicing negatively regulates virus- triggered IFN-beta signalling pathway by disrupting
44993	TBK1	18977754	DDX58 and MAVS, and inhibiting IFN-beta signalling pathways. TBK1 and IKBKE (IKKi) kinases are required for innate immune activation by B-DNA, which may be
44993	TBK1	16286919	important in antiviral innate immunity and other DNA- associated immune disorders. TBK1 and IKBKE have a pivotal role in coordinating
44993	TBK1	12692549	the activation of IRF3 and NF-kappaB in the innate immune response.

TBK1 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA and in the subsequent induction of type I IFNs. Mice lacking Tbk1 are resistant to otherwise lethal cerebral malaria. (Demonstrated in mouse)

TBK1 is a key regulator of immunological autophagy and is responsible for autophagosome maturation into bactericidal organelles. (Demonstrated in mouse) Herpes simplex virus 1 protein, UL36 ubiquitin-

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specific protease (UL36USP), deubiquitinates TRAF3 and inhibits recruitment of TBK1 and counteracts the IFNB pathway

> USP2 deubiquitinates K63-linked polyubiquitin chains from TBK1 to terminate TBK1 activation and negatively regulate IFNB1 signalling and antiviral immune response.

Endotoxin tolerance re-programs TLR4 signalling via suppression of PELI1, a positive regulator of MyD88and TIR domain-containing adapter inducing IFN-1<sup>2</sup> (TRIF)-dependent signalling that promotes K63-linked polyubiquitination of IRAK1, TBK1, and TAK1.

HIV-1 accessory proteins Vpr and Vif bind to TBK1, inhibit its autophosphorylation, and prevent induction of type I and III interferon in myeloid cells.

> Viral interferon regulatory factor 1 (vIRF1), targets TMEM173 by preventing it from interacting with TBK1, thereby inhibiting TMEM173's phosphorylation and concomitant activation, resulting in an inhibition of the DNA sensing pathway.

NF-ΰB-mediated degradation of the coactivator NRIP1 (RIP140) regulates inflammatory responses and contributes to endotoxin tolerance.

Nrip1 is degraded by the NF-kB pathway to inactivate inflammatory gene expression and promotes endotoxin tolerance.

Nrip1 plays dual roles in regulating the M1-M2 phenotype switch in macrophages: in the nucleus as an M1 enhancer and in the cytosol as an M2 suppressor.

Bcl11b, a transcription factor essential in T cell lineage commitment and maintenance, is specifically expressed in progenitors committed to the group 2 innate lymphoid cells (ILC2) lineage and is required for ILC2 development.

Bcl11bBcl11b is a factor in the differentiation of group 2Bcl11b25964370innate lymphoid cells (ILC2s).

Bcl11b sustains the genetic and functional programs, as well as lineage fidelity, of mature type 2 innate lymphoid cells.

			Map3k8 is a MEK kinase that is require for the activation of MAP kinases in myeloid cells following TLR and TNF receptor stimulation. Map3k8 is critical for production of the pro-inflammatory cytokine TNF during inflammatory responses.
127807	Map3k8	21135874	Map3k8 plays a role in facilitating both innate and adaptive antiviral responses by transducing Type I
127807	Map3k8	26241898	interferon (IFN) signals and promoting expression of IFN-stimulated genes (ISGs). II33-dependent group 2 innate lymphoid cells (ILC2s) signal via Areg-Egfr in damaged epithelia to restore epithelial barrier function and maintain tissue
151194	Egfr	26243875	homeostasis. Il33-dependent group 2 innate lymphoid cells (ILC2s) signal via Areg-Egfr in damaged epithelia to restore
179096	Areg	26243875	epithelial barrier function and maintain tissue homeostasis. II33 is produced in alveolar macrophages that have been infected with Influenza A virus. The II33-II13
157638	I133	21623379	signalling axis is required for airway hyper-reactivity in asthma. Innate lymphoid cells responding to II33 mediate
157638	1133	22119406	airway hyperreactivity independently of adaptive immunity. IL33 predominantly elicits group 2 innate lymphoid cells (ILC2) responses, and IL25 simultaneously elicits phenotypically and functionally distinct ILC2
157638	1133	23960191	and multipotent progenitor type 2 cell populations at multiple tissue sites. Chitin induces IL25, IL33, and TSLP which are required to stimulate ILC2 production of IL5 and IL13. IL5 and IL13, in turn, are required for the accumulation
157638	1133	24631157	of eosinophils and alternatively activated macrophages that are associated with allergy. II33 prevents the development of experimental cerebral malaria by orchestrating a protective immune response
157638	1133	25659095	<ul><li>via type-2 innate lymphoid cells, M2 macrophages and regulatory T cells.</li><li>Cigarette smoke decreases Il1rl1 expression on group 2 innate lymphoid cells while elevating Il1rl1 expression</li></ul>
157638	1133	25786179	on macrophages and natural killer cells, thus altering II33 responsiveness within the lung to infection. Group 2 innate lymphoid cell (ILC2)-intrinsic II33 signalling and Icosl expression promote regulatory T cell accumulation, whereas the inflammatory cytokine
157638	1133	26092469	Ifng counter-regulates these effects, in part through direct effects on ILC2s. Il33-dependent group 2 innate lymphoid cells (ILC2s) signal via Areg-Egfr in damaged epithelia to restore epithelial barrier function and maintain tissue
157638	II33	26243875	homeostasis.

			Card9 is critical for full activation of innate immunity by converging signals downstream of multiple pattern recognition receptors (PRRs) and plays a pivotal role in
148496	Card9	20351059	autonomous innate host defense against tuberculosis. Card9 is largely dispensable for the innate immune
148496	Card9	24379290	response to oropharyngeal candidiasis whereas it is vital for the adaptive Th17 response. Card9-mediated activation of the innate immune system exacerbates influenza pneumonia. Card9 deficiency improves mortality with reduced
148496	Card9	26627732	deficiency improves mortality with reduced inflammatory cytokine/chemokines. Ikbkb and other IKK kinases regulate each other by an intricate network involving phosphorylation of their catalytic and regulatory (NEMO, TANK) subunits to balance their activities during innate immunity.
141021	Ikbkb	21138416	The reversible ubiquitin editing of Nlrc5 determines
141021	Ikbkb	26620909	NIrc5 ⠀ Ikbkb interaction dynamics and plays a crucial role in precisely regulating NFΰB signalling The reversible ubiquitin editing of NIrc5 determines NIrc5 â € Ikbkb interaction dynamics and plays a
182231	Nlrc5	26620909	crucial role in precisely regulating NFI°B signalling NLRC5 is nn NLR (nucleotide-binding domain and leucine-rich repeat containing family receptors) that is part of the family of pattern recognition receptors (PRRs) and is involved in immunity against
32729	NLRC5	19414032	intracellular pathogens. NLRC5 represents a molecular switch of IFN-gamma activation sequence/IFN-specific response element signalling pathways contributing to antiviral defence
32729	NLRC5	20061403	mechanisms. NLRC5 is a negative regulator that blocks two central components of the NF-kappaB and type I interferon signalling pathways and has an important role in
32729	NLRC5	20434986	homeostatic control of innate immunity. NLRC5 is dispensable for cytokine induction in virus and bacterial infections under physiologic conditions.
32729	NLRC5	21148033	(Demonstrated in murine model) The reversible ubiquitin editing of NLRC5 determines
32729	NLRC5	26620909	NLRC5†IKBKB interaction dynamics and plays a crucial role in precisely regulating NFΰB signalling Influenza A virus non-structural protein 1, NS1, physically interacts with endogenous NLRP3
107914	NLRP3	25978411	<ul><li>downregulating NLRP3 inflammasome activation as well as NF-kB, leading to a reduction in the levels of inflammatory cytokines.</li><li>RNA cleavage products, catalyzed by RNASEL, bind to DHX33 to facilitate the formation of a complex with MAVS and NLRP3 during viral infection.</li></ul>
107914	NLRP3	25816776	

107914	NLRP3	26610398	H. pylori infection induces the expression and activation of components of NLRP3 inflammasomes in neutrophils and this activation is independent of a functional type IV secretion system, TLR2 and TLR4. Staphylococcus aureus extracellular adherence protein (Eap) binds to C4B to inhibit binding of both full- length C2 and its C2b fragment, disrupting the
124824	C2	25381436	formation of the C3 proconvertase (C4b2) and significantly diminishing the extent of S. aureus opsonophagocytosis and killing by neutrophils. Complement interfering protein (CIP) of group B Streptococcus (GBS) shows high affinity toward C4B and inhibits its interaction with C2, presumably preventing the formation of the C4BC2A convertase
124824	C2	26608922	and GBS phagocytic killing in the absence of anti-GBS antibodies. Staphylococcus aureus extracellular adherence protein (Eap) binds to C4B to inhibit binding of both full-
301132	C4B	25381436	length C2 and its C2b fragment, disrupting the formation of the C3 proconvertase (C4b2) and significantly diminishing the extent of S. aureus opsonophagocytosis and killing by neutrophils. Complement interfering protein (CIP) of group B Streptococcus (GBS) shows high affinity toward C4B
301132	C4B	26608922	and inhibits its interaction with C2, presumably preventing the formation of the C4BC2A convertase and GBS phagocytic killing in the absence of anti-GBS antibodies. Ddx58 (Rig-I) is responsible for the cytosolic
135957	Ddx58	19936053	recognition of Legionella pneumophila RNA and the subsequent induction of type I IFN response. Ddx58 and Nod2 colocalize to cellular ruffles and cell- cell junctions to form a protein complex via the CARD
135957	Ddx58	21690088	domains. Ddx58 negatively regulates ligand-induced NFkB signalling mediated by Nod2, and conversely, Nod2 negatively regulates type I interferon induction by Ddx58. (Demonstrated in human) Ddx58, through the TRAIL pathway, initiates
135957	Ddx58	21695051	apoptosis in hepatocytes infected with hepatitis C Virus. HCV envelope proteins counteract the antiviral host defence by inhibiting the expression of Ddx58. (Demonstrated in human)
			Ddx58 (RIG-I) ubiquitination is inhibited by arterivirus and nairovirus deubiquitinating enzymes (DUBs), resulting in the inhibition of RIG-I-like receptor
135957	Ddx58	22072774	(RLR)-mediated innate immune signalling. Antiviral stress granules containing Ddx58 (RIG-I) and Eif2ak2 (PKR) have a critical role in viral detection
135957	Ddx58	22912779	and innate immunity. DDX58 (RIG-I) detects cytosolic Listeria monocytogenes infections by sensing secreted bacterial
135957	Ddx58	23064150	nucleic acids.

135957	Ddx58	23553835	Ddx58 (RIG-I) is a positive regulator of NF-kB signalling via binding to Nfkb1 mRNA. Ddx58 and Ifih1 are essential pattern recognition
135957	Ddx58	23966395	receptors for protection against West Nile virus infection in vivo. Ddx58 preferentially binds to coding RNA from S.
135957	Ddx58	24692634	Typhimurium during infection leading to the expression of IFN beta andthis immunostimulatory activity depends on $5\hat{a}\xi^2$ triphosphorylation of RNA. Ddx58 is the primary pattern recognition receptor (PRR) for influenza A virus (IAV), but Ifih1 is a significant contributor to the cellular defense against
135957	Ddx58	26074083	IAV. Ddx58 acts in parallel with Zbp1 in an RNA
135957	Ddx58	26146945	polymerase III-dependent manner to initiate glial responses to herpes simplex virus-1. Mir485 exhibits bispecificity, targeting Ddx58 in cells
135957	Ddx58	26645583	<ul><li>with a low abundance of H5N1 virus and viral PB1 in cells with increased amounts of the H5N1 virus.</li><li>Mir485 exhibits bispecificity, targeting Ddx58 in cells with a low abundance of H5N1 virus and viral PB1 in cells with increased amounts of the H5N1 virus.</li></ul>
221446	Mir485	26645583	MIR485 exhibits bispecificity, targeting DDX58 in cells with a low abundance of H5N1 virus and viral PB1 in cells with increased amounts of the H5N1 virus.
127079	MIR485	26645583	IL22 induced at an early stage of L. monocytogenes
			infection enhances innate immunity against L. monocytogenes in the liver by stimulating hepatocytes
92794	PLA2G2A	26644377	to produce an antimicrobial molecule, PLA2G2A IL22 plays a role in mucosal immunity where it helps
46046	IL22	18978771	constrain inflammation and protect mucosal sites. IL22 is a member of the IL-10 cytokine family that is produced by special immune cell populations and ts primary effects on target cells include its role in innate immune defence against infections, in tumourigenesis,
46046	IL22	20870448	and in inflammatory diseases. IL22 increases the TNF-alpha-dependent induction and secretion of several immune-modulatory molecules such as initial complement factors, antimicrobial peptides and chemokines in primary keratinocytes. IL22-mediated induction of innate immunity is crucial
46046	IL22	21469124	for the maintenance of epidermal integrity during infection with Candida albicans. IL22 is produced by lymphoid tissue-inducer cells where it regulates the maintenance of colonic lymphoid structures during Citrobacter rodentium infection, a mechanism that bridges the lymphotoxin pathway to mucosal epithelial defense mechanisms. (Demonstrated
46046	IL22	21874025	in mice)

			IL22 protects intestinal stem cells from inflammatory
46046	IL22	22921121	tissue damage and regulates sensitivity to graft versus host disease. Stat3 mediates protection against intestinal infection by
			inducing innate lymphoid cell derived-II22.
46046	IL22	24412612	(Demonstrated in mice) IL22 protects against and IL22RA2 aggravates liver
46046	IL22	25476703	fibrosis and cirrhosis in chronic liver infections. IFNG interferes with the IL-1/NFKBIZ axis in $\hat{1}^2$ - glucan-activated dendritic cells and promotes T cell- mediated immune responses with increased release of
46046	IL22	25474109	IFNG and IL22, and diminished production of IL17A. In alveolar epithelium, IL22 upregulates DEFB4A gene
46046	IL22	25510212	expression via STAT3. IL22 induced at an early stage of L. monocytogenes infection enhances innate immunity against L.
46046	IL22	26644377	monocytogenes in the liver by stimulating hepatocytes to produce an antimicrobial molecule, PLA2G2A Cd47 plays a protective role against disseminated candidiasis and alters pro-inflammatory and
165137	Cd47	26010544	immunosuppressive pathways known to regulate innate and T cell immunity.
165137	Cd47	27194758	Cd47 plays a role as a negative regulator in inducing protective immune responses to influenza vaccination. IFI27 restricts viral infection by recruiting an E3 ligase, SKP2, for ubiquitination and degradation of viral
16420	SKP2	27194766	protein.
			IFI27 is a mitochondrial protein and its expression
17962	IFI27	20939681	sensitizes cells to apoptotic stimuli via mitochondrial membrane destabilization. IFI27 restricts viral infection by recruiting an E3 ligase,
17962	IF127	27194766	SKP2, for ubiquitination and degradation of viral protein. Clec4n, a C-type lectin receptor, is a pattern recognition receptor critical for immune responses to fungi. Clec4n is coupled to Syk kinase and signals via Card9 to activate NFKB, which in turns induces both immute and education immune responses.
186283	Clec4n	21267996	innate and adaptive immunity.
			Clec4n is critical for the development of house dust mite (Dermatophagoides farinae) elicited eosinophilic and neutrophilic pulmonary inflammation. Clec4n was also found to be crucial for the Th2 cytokine induction
186283	Clec4n	21357742	in the lungs and re-stimulated lymph nodes. Clec4n is expressed mainly in DCs and macrophages. Clec4n recognizes alpha-mannans with its carbohydrate recognition domain and transduces signals through association with the ITAM-containing Fc receptor gamma chain, which recruits Syk and initiates the
186283	Clec4n	21677049	Card9/NFkB signalling cascade.

186283	Clec4n	27194783	Pik3cd regulates Clec4n signalling and the generation of Th2 and Th17 immunity.
205221	D:1-2 - 1	27104702	Pik3cd regulates Clec4n signalling and the generation
205331	Pik3cd	27194783	of Th2 and Th17 immunity. IFIT3 triggers host antiviral responses by bridging
81919	IFIT3	21813773	TBK1 to MAVS, and IFIT3 plays an important role in the activation of IRF3.
01919	11115	21013773	Hepatitis B virus-induced MIR146A attenuates cell-
81919	IFIT3	27210312	intrinsic anti-viral innate immunity through targeting DDX58 and IFIT3.
01717	11115	27210312	MIR146A upregulation by CXCR4 antagonist
			AMD3100 treatment or ZBTB16 silencing, decreases CXCR4 protein expression and prevents HIV-1
			infection of leukemic monocytic cell line and CD4(+)
774160	MIR146A	25705792	T lymphocytes. Hepatitis B virus-induced MIR146A attenuates cell-
		0	intrinsic anti-viral innate immunity through targeting
774160	MIR146A	27210312	DDX58 and IFIT3. TRAF3 serves as a negative regulator of the non-
			canonical NF-kappaB pathway by specifically blocking
21240	TRAF3	15708970	the activation of NF-kappaB via TRAF2/5. TRAF3 is required for type I interferon production in
			response to intracellular double-stranded RNA.
			Similarly, a direct and specific interaction of the TRAF domain of TRAF3 with the TRAF-interaction motif
			(TIM) of MAVS is required for optimal MAVS-
21240	TRAF3	16858409	mediated antiviral responses.
21240	TRAF3	16306936	TRAF3 is a major regulator of type I interferon (IFN) production and the innate antiviral response.
			TRAF3 cooperates with ZMYND11 in the regulation of Epstein-Barr virus-derived LMP1/CTAR1-induced
21240	TRAF3	20138174	NF-kappaB activation.
			TRAF3 functions downstream of multiple TNF
			receptors and receptors that induce interferon (IFN)- alpha, IFN-beta, and IFN-lambda production, including
			Toll-like receptor 3 (TLR3). TLR3-mediated immunity
			against primary infection by herpes simplex virus-1 (HSV-1) in the central nervous system is critically
21240	TRAF3	20832341	dependent on TRAF3.
			TRAF3 is a highly versatile regulator that positively controls type I interferon production, and negatively
01040		21((0)52	regulates MAP kinase activation and alternative NFkB
21240	TRAF3	21660053	signalling. Upon sensing dsRNA or dsDNA, TRAF3 interacts
21240		22702062	with ER-to-Golgi transport proteins to induce MAVS-
21240	TRAF3	22792062	associated innate immune responses. Herpes simplex virus 1 protein, UL36 ubiquitin-
			specific protease (UL36USP), deubiquitinates TRAF3
21240	TRAF3	23986588	and inhibits recruitment of TBK1 and counteracts the IFNB pathway
21240		07012420	HACE1 plays an inhibitory role in virus-induced
21240	TRAF3	27213432	signalling by disrupting the MAVS-TRAF3 complex.

94750 131590	HACE1	27213432 19362023	HACE1 plays an inhibitory role in virus-induced signalling by disrupting the MAVS-TRAF3 complex. Casp1 is involved in key innate and healing responses to influenza A virus where Casp1(-/-) mice exhibited increased morbidity after infection with a pathogenic influenza A virus correlating with decreased neutrophil and monocyte recruitment and reduced II1b (IL-1-beta), II18 (IL-18), Tnf (TNF-alpha), II6 (IL-6), Cxcl1 (KC), and Cxcl2 (MIP-2) production.
131390	Casp1	19302023	Casp1 is part of the inflammasome complex, along with pathogen-specific nucleotide oligomerization and binding domain (NOD)-like receptors (NLRs) and in some cases the scaffolding protein ASC. Formation of the membrane-associated inflammasome complex in murine macrophages, results in cleavage of cytosolic Casp1 substrates and cell death.
131590	Casp1	19124602	
131590	Casp1	21439959	Casp1 is a component of the inflammasome and is required for inflammation in acute pancreatitis. Casp1-dependent inflammatory cell death, or pyroptosis, is only induced by viable, but not heat- killed, E. coli.
131590	Casp1	21602824	Naturally occurring variants of Casp1 differ considerably in structure and the ability to activate II1b. (Demonstrated in human)
131590	Casp1	22833538	Activation of the Nlrp3 inflammasome is detrimental during leishmaniasis. Mice lacking the inflammasome components Nlrp3, Pycard, Casp1 exhibit defective Il1b and Il18 production at the infection site and are
131590	Casp1	25689249	resistant to cutaneous Leishmania major infection. Type 1 regulatory (Tr1) cells suppress II1b transcription and Casp1 activation via an IL10R
131590	Casp1	26056255	a€ dependent mechanism Uropathogenic Escherichia coli protein TcpC attenuates activation of the Nlrp3 inflammasome by
131590	Casp1	27214553	binding both Nlrp3 and Casp1.
164567	Retnla	23355735	Retnla (RELMalpha) has a proinflammatory role in bacterial-induced colitis. Expression of Cxcl13, Ifngr1, Retnla and Mrc1 distinguishes between large and small resident
164567	Retnla	27220602	peritoneal macrophage subsets. Expression of Cxcl13, Ifngr1, Retnla and Mrc1
135936	Ifngr1	27220602	distinguishes between large and small resident peritoneal macrophage subsets. Expression of Cxcl13, Ifngr1, Retnla and Mrc1 distinguishes between large and small resident
181227	Cxcl13	27220602	distinguishes between large and small resident peritoneal macrophage subsets.

138187	Mrc1	27220602	Expression of Cxcl13, Ifngr1, Retnla and Mrc1 distinguishes between large and small resident peritoneal macrophage subsets. NFKBIA is a common component of the
4758	NFKBIA	9891086	heterogeneous IKK complex that mediates an essential step of the NF-kappaB signal transduction cascade by acting as an inhibitor of NF-kappaB. Tyrosine phosphorylation of NFKBIA is c-Src- dependent, leading to the subsequent activation of NF-
4758	NFKBIA	12429743	kappaB. Nuclear transport of the NFKBIA : RELA complex,
4758	NFKBIA	16931600	required for the appropriate regulation of NF-kappaB signalling, is facilitated by 14-3-3 proteins. NFKBIA ubiquitination and degradation is inhibited by ChlaDuhl
4758	NFKBIA	18503636	ChlaDub1, a protein of Chlamydia trachomatis, suppressing NF-kappaB activation as a result. NFKBIA degradation occurs through the TNF- stimulated formation of autophagosomes in epithelial
4758	NFKBIA	21454695	cells, which results in the prolonged activation of NFKB activity. Polymorphisms in the NFKBIA promoter are associated with pediatric lung diseases, including
4758	NFKBIA	23487427	childhood asthma, bronchiolitis and bronchopulmonary dysplasia. Haploinsufficiency of A20 (HA20) is caused by high- penetrance loss-of-function germline mutations in
4758	NFKBIA	26642243	TNFAIP3 with increased degradation of NFKBIA, nuclear translocation of RELA, increased expression of NF Î ° B mediated proinflammatory cytokines, and defective deubiquitinating activity.
			TNFAIP3 restricts TLR signals by restricting ubiquitination of TRAF6 and restricts MyD88- independent TLR signals by inhibiting Toll/interleukin 1 receptor domain-containing adaptor inducing
			interferon (IFN) beta (TICAM1)-dependent nuclear factor kappaB signals but not IFN response factor 3
97033	TNFAIP3	18268035	signalling. TNFAIP3 restricts NOD2 triggered signals by
97033	TNFAIP3	18342009	deubiquitinating RIPK2. TNFAIP3 negatively regulates the RIG-I antiviral state
97033	TNFAIP3	16306043	by blocking IRF- and NF-kappaB-mediated gene expression. TNFAIP3 negatively regulates BCL10- and CARMA3- mediated activation of NF-kappaB by means of its
			deubiquitylation activity and preventing assembly of
97033	TNFAIP3	18349075	the complex containing CARMA3, BCL10 and IKBKG (NEMO). TNFAIP3 removes lysine-63 linked ubiquitin chains from RIPK1 and polyubiquitinates RIPK1 with K48- linked ubiquitin chains, targeting it for proteosomal
07022		15759507	degradation, and hence down-regulating NF-kappaB
97033	TNFAIP3	15258597	signalling pathway.

97033	TNFAIP3	17709380	TNFAIP3 is phosphorylated by IKBKB and this increases the ability of TNFAIP3 to inhibit the NF- kappaB signalling pathway. TNFAIP3 is an early NF-kappaB-responsive gene that encodes a ubiquitin-editing protein that is involved in the negative feedback regulation of NF-kappaB
97033	TNFAIP3	19008218	signalling and thus is a central gatekeeper in inflammation and immunity. TNFAIP3 (A20) and TAX1BP1 inhibit antiviral
97033	TNFAIP3	20304918	signaling by targeting TBK1/IKKi kinases and disrupting a TRAF3-TBK1-IKKi signalling complex. TNFAIP3 and TRAF6 differentially regulate TLR4- induced autophagy during inflammatory responses by modulating K63-linked ubiquitination of BECN1
97033	TNFAIP3	20798608	(Beclin 1). TNFAIP3 is a deubiquitinase that counteracts E3
97033	TNFAIP3	21119682	ligases and therefore play a prominent role in the down-regulation of NF-ΰB signalling and homeostasis. TNFAIP3 expression and TNFAIP3-IRAK1interaction are important for endotoxin tolerance. TNFAIP3 over- expression inhibits LPS-induced activation of NFKB, and is mechanistically linked to endotoxin tolerance through the reprogramming of TLR4 signalling.
97033	TNFAIP3	21220427	TNFAIP3 promotes intestinal epithelial barrier integrity and inhibits LPS-induced loss of the tight
97033	TNFAIP3	22031828	junction protein occludin. (Demonstrated in mice) Haploinsufficiency of A20 (HA20) is caused by high- penetrance loss-of-function germline mutations in TNFAIP3 with increased degradation of NFKBIA, nuclear translocation of RELA, increased expression of NF Î ° B mediated proinflammatory cytokines, and
97033	TNFAIP3	26642243	defective deubiquitinating activity. Ifih1 is an RNA helicase and is a key component in activating the expression of type I IFN in response to viral infection. Viral mRNA with 5' cap and 3' poly(A) from parainfluenza virus 5 is able to activate IFN expression through Rnasel-Ifih1 signalling pathway.
174129	Ifih1	21245317	Ifih1 (MDA5) is responsible for the cytosolic
174129	Ifih1	19936053	recognition of Legionella pneumophila RNA and the subsequent induction of type I IFN response. Ifih1 deficiency results in a delayed type I IFN and attenuated type III IFN response to rhinovirus infection, leading to a transient increase in viral titer. Upon recognition of viral dsRNA, Ifih1 synergizes with Tlr3 to induce pro-inflammatory signals leading
174129	Ifih1	21637773	to airways inflammation and hyper-responsiveness. Paramyxovirus V proteins bind to IFIH1 (MDA5) to disrupt viral RNA recognition and induction of
174129	Ifih1	23328395	antiviral immunity.

174129	Ifih1	23966395	Ddx58 and Ifih1 are essential pattern recognition receptors for protection against West Nile virus infection in vivo.
			Plasmodium RNA is a pathogen-associated molecular pattern (PAMP) capable of activating a type I IFN response via the cytosolic pattern recognition receptors
174129	Ifih1	24362933	Ifih1 and Mavs, as well as via transcription factors Irf3 and Irf7. Arl5b negatively regulates the antiviral innate immune
174129	Ifih1	25451939	response by binding to Ifih1 and prevents the subsequent interaction of Ifih1 to poly(I:C). Ddx58 is the primary pattern recognition receptor (PRR) for influenza A virus (IAV), but Ifih1 is a
174129	Ifih1	26074083	significant contributor to the cellular defense against IAV. Transgenic picornavirus RNA-dependent RNA
174129	Ifihl	26633895	polymerase (RdRP) expression in mice produces a quantitatively dramatic, sustained, effective antiviral interferon-stimulated genes (ISG) network, which requires the MDA5-MAVS pathway.
174129	11111	20033893	Selective loss of the histone-lysine N-methyltransferase Ezh2 (enhancer of zeste homolog 2) or inhibition of its enzymatic activity increases generation of the Il2rb
159374	Il2rb	26668377	<ul><li>(CD122) natural killer (NK) precursors and mature NK progeny.</li><li>Selective loss of the histone-lysine N-methyltransferase Ezh2 or inhibition of its enzymatic activity increases</li></ul>
143636	Ezh2	26668377	generation of the Il2rb (CD122) natural killer (NK) precursors and mature NK progeny.
192196	Ifng	23754402	<ul><li>IFNG is produced by neutrophils to mount host protection against intracellular pathogens.</li><li>IFNG produced by natural killer cells exacerbates Listeria monocytogenes infection by inhibiting</li></ul>
192196	Ifng	23818011	granulocyte recruitment. Ticam2 mediates antibacterial defence during Gram-
192196	Ifng	26065469	negative pneumonia by inducing Ifng at the primary site of infection. Group 2 innate lymphoid cell (ILC2)-intrinsic II33 signalling and Icosl expression promote regulatory T cell accumulation, whereas the inflammatory cytokine
192196	Ifng	26092469	Ifng counter-regulated the effects of II33, in part through direct effects on ILC2s. Irf8-Ifng circuit is a novel gastric innate immune
192196	Ifng	26843324	mechanism in the host defense against infection with Helicobacter pylori. De-SUMOylation of IRF8 at residue Lys310 acts as a
196509	Irf8	22942423	molecular mechanism to trigger innate immune responses in activated macrophages. Irf8-Ifng circuit is a novel gastric innate immune
196509	Irf8	26843324	mechanism in the host defense against infection with Helicobacter pylori.

193230	Sppl3	26851218	Sppl3 is an intramembrane aspartyl protease that controls natural killer cell maturation and cytotoxicity. Ahr deficiency impairs TLR and NFkB-mediated
134957	Ahr	21683686	proinflammatory gene expression after activation by a classical stimulus, such as LPS. Innate expression of Ahr plays a protective role in T-
134957	Ahr	23954130	cell-induced colitis by suppressing T helper 17 cells, thus inhibiting proinflammatory cytokine production. Uropathogenic Escherichia coli suppresses neutrophil migration early in bacterial cystitis by eliciting an Ido1-mediated increase in local production of
134957	Ahr	26857571	kynurenines, which act through the Ahr to impair neutrophil chemotaxis. Ido1 limits innate and adaptive immunity to apoptotic self-antigens. Ido1-mediated inhibition of inflammation plays a key role in suppressing systemic
142448	Ido 1	22355111	autoimmune diseases. Uropathogenic Escherichia coli suppresses neutrophil migration early in bacterial cystitis by eliciting an Ido1-mediated increase in local production of kynurenines, which act through the Ahr to impair
142448	Ido1	26857571	neutrophil chemotaxis.
154384	Lgals3	22486577	Lgals3 influences the course of malaria in a Plasmodium species-specific manner.
154384	Lgals3	26857579	Lgals3 plays an important role in innate immunity to infection and colonization of Helicobacter pylori. Tuft cells express II25 and elicit group 2 innate
212683	Trpm5	26847546	lymphoid cells in a Trpm5-dependent manner in response to parasite colonization. Chitin induces IL25, IL33, and TSLP which are required to stimulate ILC2 production of IL5 and IL13. IL5 and IL13, in turn, are required for the accumulation of eosinophils and alternatively activated macrophages
162540	1125	24631157	that are associated with allergy. Tuft cells express II25 and elicit group 2 innate lymphoid cells in a Trpm5-dependent manner in
162540	1125	26847546	response to parasite colonization. Gata3 plays a generalized role in innate lymphoid cell (ILC) lineage determination and is critical for the development of gut Rorc+ group 3 ILCs subsets that
169968	Rorc	24419270	maintain mucosal barrier homeostasis. Retinoic acid (RA) induces the maturation of lymphoid tissue inducer cells in developing lymph nodes by inducing RA receptor binding to the promoter region of
169968	Rorc	24670648	Rorc.
169968	Rorc	26878233	Rorc is differentially required in the maintenance of TH17 cell and group 3 innate lymphoid cell responses.

Aberrant mitochondrial DNA (mtDNA) packaging promotes escape of mtDNA into the cytosol, where it engages the DNA sensor Mb21d1 and promotes Tmem173-Irf3-dependent signalling to elevate IFN-stimulated gene expression, potentiate type I IFN responses and confer broad viral resistance.

The cationic polymer and vaccine adjuvant chitosan can engage the Tmem173/Mb21d1 (STING/cGAS) pathway to trigger innate and adaptive immune responses.

Tmem173 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA and in the subsequent induction of type I IFNs.

Tmem173Tmem173 activatesStat6 during viral infection to<br/>induce genes responsible for immune cell homing.

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Dengue viral NS2B3 protease complex cannot degrade murine TMEM173, which confers protection against the viral infection.

Cyclic dinucleotides initiate the production of Tmem173(STING)-dependent proinflammatory genes and a negative-feedback to prevent sustained production that may otherwise lead to inflammation.

24131791 Cyclic-di-GMP-induced levels of Ifi202b suppress the expression of Tmem173 (STING).

The innate immune system plays a role in immunogenic tumour recognition. Tumor-cell-derived DNA triggers Ifnb1 production and dendritic cell activation via Tmem173 and Irf3 cytosolic DNA sensing pathways.

Unrepaired DNA lesions induce type I interferons via the Tmem173 pathway, resulting in enhanced anti-viral and anti-bacterial responses in Atm (-/-) mice.

Tmem173-deficient macrophages fail to express negative regulators of immune activation and are hyperresponsive to TLR ligands, producing abnormally high levels of proinflammatory cytokines.

Aberrant mitochondrial DNA (mtDNA) packaging promotes escape of mtDNA into the cytosol, where it engages the DNA sensor Mb21d1 and promotes Tmem173-Irf3-dependent signalling to elevate IFN-stimulated gene expression, potentiate type I IFN responses and confer broad viral resistance.

26590319 DNA vaccine-induced, Irf7-dependent signalling, as part of the Tmem173 (Sting) pathway, is critical for generation of both innate cytokine signalling and antigen-specific B and T cell responses.

The cationic polymer and vaccine adjuvant chitosan<br/>can engage the Tmem173/Mb21d1 (STING/cGAS)<br/>pathway to trigger innate and adaptive immune<br/>responses.Tmem17326944200

106840	TLR5	15069060	TLR5 engagement with its ligand, flagellin, results in the activation of MAP kinases (ERK1/2, JNK, and p38) and degradation of NFKBIA (I-kappa-B-alpha). Toll-like receptor 5 (TLR5) is the TLR activated by bacterial flagellin and TLR5 residue 268 is responsible
106840	TLR5	17283206	for human and mouse discrimination between flagellin molecules. TLR5 recognizes bacterial flagellin from both Gram- positive and Gram-negative bacteria, and its activation
106840	TLR5	11323673	mobilizes the nuclear factor NF-kappaB and stimulates tumour necrosis factor-alpha (TNF) production. TLR5 expressed on the basolateral surface of intestinal epithelia mediates innate immune inflammatory responses to Salmonella by inducing epithelial pro-
106840	TLR5	11489966	inflammatory gene expression. TLR5 forms heteromeric complexes with TLR4 as well as homomeric complexes. Signaling via heteromeric TLR5/TLR4 complexes is involved in the induction of macrophage nitric oxide (NO) production by Gram-
106840	TLR5	12794153	negative flagellin. TLR5 forms a dimer upon binding to bacterial
106840	TLR5	22434932	flagellin. TLR5 is highly expressed in mucosal dendritic cells and TLR5 signalling restricts regulatory T cell
106840	TLR5	22545147	generation. (Demonstrated in mice) TLR5 deficiency in mice leads to a transient inability
106840	TLR5	22863420	to manage proteobacteria which promotes chronic gut inflammation. (Demonstrated in mouse) CXCR1 regulates anti-Pseudomonas neutrophil responses through modulation of reactive oxygen species and interference with TLR5 expression.
106840	TLR5	26950764	CXCR1 regulates anti-Pseudomonas neutrophil responses through modulation of reactive oxygen species and interference with TLR5 expression.
80821	CXCR1	26950764	Stat6 is phosphorylated upon viral infection and
195602	Stat6	22000020	translocates to the nucleus to induce genes responsible for immune cell homing.
195602	Stat6	26953325	Stat6 signalling negatively regulates Î <sup>3</sup> Î′17 T cells, which plays a front-line role in mucosal immunity. Mir342 is involved in a novel post-transcriptional viral defence mechanism in IFN-activated macrophages by directly targeting Srebf2, Idi1, Dhcr7 and Sc4mol of
212845	Dhcr7	26938778	the sterol pathway. Mir342 is involved in a novel post-transcriptional viral defence mechanism in IFN-activated macrophages by directly targeting Srebf2, Idi1, Dhcr7 and Sc4mol of
158311	Sc4mol	26938778	the sterol pathway.

			Mir342 is involved in a novel post-transcriptional viral defence mechanism in IFN-activated macrophages by directly targeting Srebf2, Idi1, Dhcr7 and Sc4mol of
129163	Idi 1	26938778	the sterol pathway. Srebf2 is a key transcriptional regulator of sterol
			biosynthesis in lipid metabolism, and Srebp2 protein levels in macrophages are negatively regulated by type
166203	Srebf2	21408089	I interferon signalling during viral infection.
			Mir342 is involved in a novel post-transcriptional viral defence mechanism in IFN-activated macrophages by
166000	G 1 00	2(020770	directly targeting Srebf2, Idi1, Dhcr7 and Sc4mol of
166203	Srebf2	26938778	the sterol pathway. Mir342 is involved in a novel post-transcriptional viral
			defence mechanism in IFN-activated macrophages by
221078	Mir342	26938778	directly targeting Srebf2, Idi1, Dhcr7 and Sc4mol of the sterol pathway.
	-		Epithelial cell-derived cytokine Tgfb1 has a central role
			in the generation of the pulmonary immune response by enhancing the chemoactivity of type 2 innate
158292	Tgfb1	26588780	lymphoid cells.
			Sod1 is essential in protecting hepatocytes from virus- induced, interferon-driven oxidative damage in the
174662	Sod1	26588782	liver.
			LGALS3 exerts a regulatory role in innate immunity by diminishing IL-1beta production and thus affecting
7247	LGALS3	18825751	resistance to Rhodococcus equi infection.
			LGALS3, an abundant protein in macrophages and epithelial cells, belongs to a family of beta-galactoside-
			binding proteins, the galectins, with many proposed
7247	LGALS3	19951367	functions in immune response, development, differentiation, cancer and infection.
/24/	LUALSS	19931307	LGALS3 is part of the galectin family of proteins that
			have emerged as autonomous bacteria-killing agents,
7247	LGALS3	20208507	pointing to a principal role of these proteins in innate immunity.
			LGALS3 influences the course of malaria in a
7247	LGALS3	22486577	Plasmodium species-specific manner. (Demonstrated in mice)
			LGALS1 and LGALS3 play opposing roles in the
7247	LGALS3	26589797	inflammatory responses to Trichomonas vaginalis infection.
			LGALS1, the prototype of a family of beta-galactoside-
			binding proteins, is involved in monocyte chemoattraction at sites of inflammation where it
(0.5.2			stimulates monocyte migration in a dose-dependent
6929	LGALS1	19561030	manner via the p44/42 MAP kinase pathway. LGALS1 and LGALS3 play opposing roles in the
			inflammatory responses to Trichomonas vaginalis
6929	LGALS1	26589797	infection.

205873	Aim2	20401524	Aim2 is uniquely involved in sensing infection with the intracellular bacteria Francisella tularensis and subsequently triggering caspase-1-mediated pro- inflammatory cytokine production and macrophage cell death, which activate other components of the immune system and eliminate the infected macrophages. Aim2 is required for innate immune recognition of Francisella tularensis where Aim2-deficient mice display an increased susceptibility to F. tularensis
205873	Aim2	20457908	infection compared with wild-type mice. Aim2-containing inflammasomes are activated in response to cytosolic DNA; this response is augmented in keratinocytes from psoriatic lesions and contributes to the auto-inflammatory disease. (Demonstrated in human)
205873	Aim2	21562230	During influenza A virus infection, host-derived DNA accumulates in the lung microenvironment and is sensed by Aim2, which limits immune-mediated damage to infected tissues.
205873	Aim2	26590313	Irf7 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA and in the subsequent induction of type I IFNs. Mice lacking
212081	Irf7	21820332	Irf3/Irf7 are resistant to otherwise lethal cerebral malaria. Plasmodium RNA is a pathogen-associated molecular pattern (PAMP) capable of activating a type I IFN response via the cytosolic pattern recognition receptors
212081	Irf7	24362933	Ifih1 and Mavs, as well as via transcription factors Irf3 and Irf7. Genetic deletion of Eif4ebp1 or Eif4ebp2 potentiates
212081	Irf7	25531441	innate antiviral immunity by enhancing translation of Irf7. DNA vaccine-induced, Irf7-dependent signalling, as part of the Tmem173 (Sting) pathway, is critical for
212081	Irf7	26590319	generation of both innate cytokine signalling and antigen-specific B and T cell responses. Cxcr3 expression on recruited peritoneal macrophages and granulocytes increases following sepsis, and deletion of Cxcr3 significantly increases mortality to a
164788	Cxcr3	21518789	septic challenge in neonatal mice.

Inflammatory monocyte activation status, as measured by dual production of TNF-alpha and IL-12, was severely impaired in Cxcr3(-/-) mice. Deletion of Cxcr3 in mice resulted in selective loss of ability to control T. gondii infection specifically in the lamina propria compartment.

164788	Cxcr3	24130498	compartment.
107700	CAUS	24130470	Cxcr3 expression in innate CD8+ T cells defines
164788	Cxcr3	25466888	protective antibacterial and cancer immunity upon Il15 stimulation.
			A Cxcr3-dependent innate antiviral pathway operates at
164700	<b>C 2</b>	26505800	epithelial surfaces to induce chemokines and neutrophil
164788	Cxcr3	26595890	activity prior to the induction of interferons. Invariant natural killer T cell activation induced by
			mast cells exposed to alpha-galactosylceramide is
204791	Cd48	26564814	regulated by costimulatory molecules Cd48 and Tnfsf4.
			Invariant natural killer T cell activation induced by mast cells exposed to alpha-galactosylceramide is
201525	Tnfsf4	26564814	regulated by costimulatory molecules Cd48 and Tnfsf4.
			Clec4e, a C-type lectin receptor, is a pattern recognition
			receptor critical for immune responses to fungi. Clec4e is coupled to Syk kinase and signals via Card9 to
			activate NFKB, which in turns induces both innate and
			adaptive immunity.
186402	Clec4e	21267996	The expression of Clec4e (Mincle) and its downstream
			signal phospho-Syk/Syk increases after cerebral
186402	Clec4e	24212132	ischemia and reperfusion.
			Clec4d is an inducible myeloid-expressed C-type lectin
186402	Clec4e	26558717	receptor, whose expression is tightly linked to that of Clec4e.
			Clec4d is an inducible myeloid-expressed C-type lectin
186345	Clec4d	26558717	receptor, whose expression is tightly linked to that of Clec4e.
180545	Clec4u	20338717	Map1s (Mtap1s) controls bacterial phagocytosis
			through TLR signalling by interacting directly with
164802	Mtap1s	26565030	Myd88.
			Illb acts as a growth factor for neutrophil progenitors and as a survival factor for mature neutrophils. In the
			absence of Ikbkb, the Il1b production is enhanced and
			provides a compensatory mechanism for maintaining
205936	II1b	21170027	antibacterial defense when NFKB is inhibited.
			Illb secretion in macrophages is regulated by
			autophagy by two mechanisms; sequestering of pro- Illb in autophagosome during TLR stimulation, and
			processing/secretion of II1b in a NIrp3- and TRIF-
			dependent manner.
205936	II1b	21228274	

205936	Il1b	21602824	Il1b secretion is induced only during viable E. coli infection (as oppose to heat-killed E. coli or LPS), viable bacteria specifically elicit cleavage of pro-Il1b.
203930	1110	21002824	Il1b derived from alveolar macrophages is the critical mediator which induces chemokine production in nonhematopoietic cells in the lung, resulting in swift and robust recruitment of infection-controlling
205936	II1b	21270399	neutrophils into the airways. Il1b secretion is tightly regulated by the redox status in myeloid cells. TLR engagement in monocytes induces ROS generation followed by a sustained antioxidant response and efficient Il1b secretion. In macrophages, the antioxidant systems are in an upregulated state, and therefore buffers the TLR induction of the redox response, which results in low Il1b processing and
205936	II1b	21628463	secretion. (Demonstrated in human) Il1b is an inflammatory cytokine that binds to its primary receptor, Il1r1, that then recruits the accessory protein Il1rap to form a signalling-competent
205936	Il1b	22426547	heterotrimeric complex. (Demonstrated in human) Under acidic conditions both pro-inflammatory forms of Il1a and Il1b are regulated independently of the
205936	Il1b	24022484	NLRP3 inflammasome. Group B streptococcus induces II1b, and activates the NLRP3 inflammasome by a mechanism that requires hemolysin-mediated lysosomal leakage, which enhances the interaction of bacterial RNA with
205936	II1b	24692555	NLRP3. Actin polymerization is required for Nlrc4-dependent regulation of intracellular bacterial burden, inflammasome assembly, pyroptosis, and Illb
205936	Il1b	25422455	production. Autophagy causes PELI3 degradation during Tlr4- signalling, subsequently inhibiting Il1b expression and
205936	II1b	25483963	impairing the hyperinflammatory phase during sepsis. Activation of the Nlrp3 inflammasome is detrimental during leishmaniasis. Mice lacking the inflammasome components Nlrp3, Pycard, Casp1 exhibit defective Il1b and Il18 production at the infection site and are
205936	II1b	25689249	resistant to cutaneous Leishmania major infection. Escherichia coli toxin CNF1 promotes the maturation/secretion of Il1b while the $\hat{1} \pm$ -hemolysin toxin inhibits Il1b secretion without affecting the
205936	II1b	25781937	recruitment of Ly6g+ cells. Mirlet7f and its target Tnfaip3 regulate immune responses to Mycobacterium tuberculosis and control bacterial burden by augmenting the production of Tnf
205936	Il1b	25683052	and II1b.

			Defb1 is important for the control of early mucosal Candida infection and plays a critical role in the induction of innate inflammatory mediators including,
205936	II1b	25595775	Il1b, Il6, Cxcl1, Il17a, and Il17f. Type 1 regulatory (Tr1) cells suppress Il1b transcription and Casp1 activation via an IL10R
205936	II1b	26056255	<ul> <li>†dependent mechanism.</li> <li>NLRP3 inflammasome formation is dispensable for alum-induced innate immunity but Il1a and Il1b are both necessary for alum-induced neutrophil influx in</li> </ul>
205936	Il1b	26536497	vivo. Ill8 secreted by inflammatory monocytes is critical for the differentiation of CD8(+) T and NK lymphocytes
163502	1118	22940097	into antimicrobial effector cells. Both Nlrp3 and Nlrp1a are important regulators of Toxoplasma proliferation and Il18 signaling is
163502	1118	24549849	required to mediate host resistance to acute toxoplasmosis. Flagellin induces Tlr5-dependent Il22 production and Nlrc4-dependent Il18 production to promote a
163502	II18	25395539	protective gene expression program in intestinal epithelial cells and elimination of rotavirus-infected cells. Activation of the Nlrp3 inflammasome is detrimental
163502	1118	25689249	Activation of the Nifp's inflammasome is detrimental during leishmaniasis. Mice lacking the inflammasome components Nlrp3, Pycard, Casp1 exhibit defective Il1b and Il18 production at the infection site and are resistant to cutaneous Leishmania major infection. Il22 augments the expression of Il18 mRNA and inactive precursor protein (proIL-18) in intestinal epithelial cells after Toxoplasma gondii or Citrobacter
163502	1118	25680273	rodentium infection and maintains the homeostatic amount of proIL-18 in the ileum. Microbiota-associated metabolites modulate Nlrp6
163502	II18	26638072	inflammasome signalling, epithelial II18 secretion, and anti-microbial pathways.
163502	II18	26638073	Colitis severity is controlled at the level of II18 signalling in intestinal epithelial cells. Nlrp6 is a negative regulator of inflammatory
211907	Nlrp6	22763455	signalling and impedes the clearance of both Gram- positive and -negative bacterial pathogens. Nlrp6 functions with Dhx15 as a viral RNA sensor to
211907	Nlrp6	26494172	induce IFN-stimulated genes, and this effect is especially important in the intestinal tract. Microbiota-associated metabolites modulate Nlrp6
211907	Nlrp6	26638072	inflammasome signalling, epithelial II18 secretion, and anti-microbial pathways.

Atg12::Atg5 conjugate is a key regulator of the autophagic process to eliminate pathogens such as Streptococcus, M. tuberculosis, Listeria, and herpesvirus. Atg12::Atg5 also associates with components of the RIG-I pathway to negatively regulate type I IFN response and promote RNA virus replication.

148568	Atg5	17921696	reprioution.
140500	Alg.	17921090	Atg5 knockout mice develop systemic and hepatic inflammation with high-fat diet and low-dose lipolysaccharide treatment.
148568	Atg5	25650776	
148568	Atg5	26649827	Atg5 plays a unique role in protection against M. tuberculosis by preventing polymorphic mononuclear cell (PMN)-mediated immunopathology. Loss of Atg5 in PMNs can cause susceptibility to M. tuberculosis. IL10 expression is regulated in different immune cells
106270	IL10	20154735	has revealed some of the molecular mechanisms involved at the levels of signal transduction, epigenetics, transcription factor binding and gene activation.
			IL10 is a potent anti-inflammatory cytokine that is crucial for down-regulating pro-inflammatory genes which are induced by Toll-like Receptor (TLR) signalling. It also plays a role in microRNA function, specifically its inhibitory effect on miR-155 expression
106270	IL10	20435894	in response to LPS. IL10 is a pleiotropic cytokine released in many tissues that mediates anti-inflammatory effects. IL10 also has an immunomodulatory role by stimulating NKG2D ligand expression on macrophages, thereby rendering
106270	IL10	20883317	them susceptible to natural killer (NK) cell elimination. IL10 contributes to antiviral innate immunity during acute infection by restricting activation-induced death in natural killer (NK) cells. Blockade of II10 receptor during acute murine cytomegalovirus (CMV) infection markedly reduced the accumulation of cytotoxic NK cells in the spleen and lung. (Demonstrated in murine
106270	IL10	21849677	model) IL10 has opposing functions in anti-microbial responses in its capacity to mediate protective immunity against some organisms but increase
106270	IL10	22268692	susceptibility to other infections. IL10-mediated suppression of natural killer/dendritic cell crosstalk leads to prolonged mouse cytomegalovirus (MCMV) persistence due to poor priming of MCMV-specific T cells. (Demonstrated in
106270	IL10	22876184	mouse) IL10 induces MIR187 to limit the expression of pro-
106270	IL10	23071313	inflammatory cytokines. MIR145 directly targets HDAC11 to promote IL10
106270	IL10	23980205	expression in TLR4-triggered macrophages.

			HIV-1 infection of macrophages modulates host responses to co-infection with Mycobacterium tuberculosis by attenuating IL10 responses thus contributing to the pathogenesis of tuberculosis in
106270	IL10	24265436	HIV-1 $\hat{a}\in$ infected patients. Secreted CCNA2 (CCN1) promotes anti-inflammatory cytokine IL10 release from epithelial cells via integrin $\hat{1}\pm V\hat{1}^2$ 6-PKC, and this subsequently suppresses TNF,
106270	IL10	25005359	CXCL2 and neutrophil infiltration in the lungs. MIR29A inhibits IL10-induced cytokine release by
106270	IL10	26535690	targeting JAK-STAT3 in monocytes during sepsis. STAT3 is a transcription factor that mediates
50702	STAT3	17971840	interleukin-10 (IL-10) cytokine signalling. STAT3 and SRC have a role in the immunoregulation of dendritic cells (DCs) where apoptotic cells-induced inhibition of dendritic cells (DCs) requires MerTK-
50702	STAT3	19667404	dependent activation of SRC and STAT3. STAT3 mediates mucosa-protective and anti-
50702	STAT3	20040863	inflammatory functions in epithelial and myeloid cells and promotes inflammation in T cells. STAT3 binds to multiple genes involved in Th17 cell differentiation, cell activation, proliferation, and survival, regulating both expression, epigenetic
50702	STAT3	20493732	modifications, and targets the T cell function in inflammation and homeostasis. STAT3 is an essential mediator of emergency granulopoiesis via its regulation of transcription factors CEBPA and CEBPB that direct granulocyte colony- tional time factors (C CSE) remember much id
50702	STAT3	20581311	stimulating factor (G-CSF)-responsive myeloid progenitor expansion. STAT3 is a negative regulator of type I IFN-mediated
50702	STAT3	21810606	anti-viral responses. (Demonstrated in mouse) In alveolar epithelium, IL22 upregulates DEFB4A gene
50702	STAT3	25510212	expression via STAT3. MIR29A inhibits IL10-induced cytokine release by
50702	STAT3	26535690	targeting JAK-STAT3 in monocytes during sepsis. MIR29A inhibits IL10-induced cytokine release by
771638	MIR29A	26535690	targeting JAK-STAT3 in monocytes during sepsis. Trim12c interacts with Traf6, a key protein in pathogen recognition receptor signalling, and reciprocally
204141	Trim12c	26503954	enhances its ubiquitination, leading to cooperative activation of IFN and NF-kB pathways. MX1 is an interferon-induced member of the dynamin superfamily of large GTPases, which inhibit a wide
3941	MX1	18062906	range of viruses by blocking an early stage of the replication cycle. MX1 GTPase is a key mediator of cell-autonomous innate immunity against a broad range of viruses such
3941	MX1	20538602	as influenza and bunyaviruses. MX1 expression is upregulated by alpha-defensins in
3941	MX1	22531919	gigival epithelial cell.
			Antiviral specificity of MX1 against orthomyxoviruses (influenza A and Thogoto viruses) is determined by a
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2041	MV1	24440002	few critical amino acids in the disordered loop L4 of
3941	MX1	24448803	MX1. NRAV, a long noncoding RNA, modulates antiviral responses by negatively regulating the initial transcription of multiple critical interferon-stimulated
3941	MX1	25525793	genes, including IFITM3 and MX1, by affecting their histone modification. MX1 prevents influenza A virus (IAV) replication by disassembling into dimers and binding to IAV
3941	MX1	26507657	disassembling into dimers and binding to IAV nucleoprotein synthesized following primary transcription.
			Lyn/PI3K module negatively regulates activation of murine macrophages while Inpp5d (SHIP-1) promotes
128365	Lyn	20385881	it.
			Overexpression of Lyn results in endotoxin hypersensitivity due to the increased activation of dendritic cells leading to an over-production of Ifng by
128365	Lyn	22491248	natural killer cells. Lyn regulates the macrophage basal-state signalling
			checkpoint, and the signalling reorganization initiated by receptor clustering allows cells to discriminate
128365	Lyn	26517880	optimally between pathogens and nonpathogens.
			Nlrp6 functions with Dhx15 as a viral RNA sensor to induce IFN-stimulated genes, and this effect is
163641	Dhx15	26494172	especially important in the intestinal tract. Il17a signalling enhances the mRNA stability of
136575	Il17a	21822258	chemokine Cxcl1 through Traf3ip2, Traf2-Traf5 and the RNA-binding protein Srsf1.
			Il17a is significantly upregulated in both S. pyogenes inoculated and mock inoculated mice, indicating that
136575	Il17a	22384827	the cytokine production can be triggered by inoculation trauma alone.
			IL17A inhibits the release of IL23 during pulmonary inflammation and serves as part of negative feedback
136575	Il17a	23295184	loop to control antibacterial immunity. Il17a is an intrinsic regulator in coordinating neutrophil
126575	1117	0.400.777.47	and macrophage antimicrobial activity to provide
136575	Il17a	24337746	protection against acute pneumonic plague. In mouse model of leukocyte adhesion deficiency type
			I disease, defective neutrophil recruitment is associated with dysregulated local overproduction of IL17, which
136575	Il17a	24670684	drives inflammatory periodontal bone loss. Defb1 is important for the control of early mucosal
			Candida infection and plays a critical role in the induction of innate inflammatory mediators including,
136575	Il17a	25595775	Il1b, Il6, Cxcl1, Il17a, and Il17f.
			Il17a is rapidly produced during lung injury and significantly contributes to early immunopathogenesis,
136575	Il17a	26488187	a process that is orchestrated largely by a distinct population of pulmonary type 3 innate lymphoid cells.

			Hyperactivated Ern1 (IRE1 $\hat{I}$ ±) increases Txnip mRNA stability by reducing levels of a Txnip destabilizing microRNA, miR-17. In turn, elevated Txnip protein activates the Nlrp3 inflammasome, causing procaspase-
213476	Ern1	22883233	1 cleavage and interleukin 1 $\hat{I}^2$ (IL-1 $\hat{I}^2$ ) secretion. (Demonstrated in human) ERN1-mediated persistent reactive oxygen species generation is a mechanism used by macrophages to kill bacterial pathogens that evade the initial oxidative burst.
213476	Ern1	26173697	C5 is the fifth component of complement and is comprised of alpha (C5a) and beta (C5b) polypeptide chains, C5a is a complement anaphylatoxin that can stimulate the generation of nitric oxide along with the secretion of histamine and leukotriene LTC4 from several cell types and can bring about an increase in vascular permeability that facilitates eosinophil
83167	C5	18039528	accumulation at sites of allergic inflammation. C5 is the fifth component of complement and is comprised of alpha (C5a) and beta (C5b) polypeptide chains. C5b first associates with complement component 6 (C6), which in turn initiates the assembly
83167	C5	10441679	of the cytolytic membrane attack complex (MAC). C5 fragments are released from neutrophils upon their activation of the alternative complement pathway and this further amplifies neutrophil pro-inflammatory
83167	C5	21063021	responses. C5 (C5a) is a critical mediator in blood during Candida
83167	C5	25539819	albicans infection. Staphylococcus aureus-induced neutrophil dysfunction correlates with the loss of C5AR1 from the neutrophil cell surface and results in C5a (C5)-induced
83167	C5	26176669	CEACAM8 overexpression. Staphylococcus aureus-induced neutrophil dysfunction correlates with the loss of C5AR1 from the neutrophil cell surface and results in C5a (C5)-induced
54589	CEACAM8	26176669	CEACAM8 overexpression. C5AR1 engage in crosstalk with TLR2 and Porphyromonas gingivalis, a major oral and systemic pathogen with complement C5 convertase-like activity, synergizes with C5a (fragment of complement protein C5) to increase cyclic adenosine monophosphate (cAMP) concentrations, resulting in suppression of macrophage immune function and enhanced pathogen
59445	C5AR1	20159852	survival.

			C5AR1 is a receptor for C5a, a chief component of complement activation produced via all three complement pathways (i.e., lectin, classical, and alternative), stimulated tissue-resident macrophages, but not dendritic cells, to produce inflammatory cytokines including IL-6, in synergy with Toll-like receptor signalling or, notably,
59445	C5AR1	20457757	granulocyte/macrophage colony-stimulating factor (GM-CSF). Increasing circulating C5AR1 serum concentration, coupled with decreased C5AR1 expression on
59445	C5AR1	23479227	neutrophils is correlated with lethality from septic shock. Staphylococcus aureus-induced neutrophil dysfunction correlates with the loss of C5AR1 from the neutrophil cell surface and results in C5a (C5)-induced
59445	C5AR1	26176669	CEACAM8 overexpression. Zbp1 interacts with Ripk3 to mediate virus-induced
213369	Zbp1	22423968	necrosis. Ddx58 acts in parallel with Zbp1 in an RNA polymerase III-dependent manner to initiate glial
213369	Zbp1	26146945	responses to herpes simplex virus-1. Inactivation of Hs2st1 in neutrophils substantially reduces their bactericidal activity, and Hs2st1-deficient
197593	Hs2st1	26150541	mice are more susceptible to systemic infection with the pathogenic bacterium group B Streptococcus. Dusp1 modulates the phosphorylation status of mBNA destabilizing protein 7fp26 to regulate
163345	Zfp36	26019272	mRNA-destabilizing protein Zfp36 to regulate macrophage immune response to lipopolysaccharide. Dusp1 is a negative regulator of MAPK-dependent induction of II6 and II8 in response to the coronavirus infectious bronchitis virus (IBV). (Demonstrated in
155234	Dusp1	21959016	human) Dusp1 antagonizes p38 MAPK activity to induce II12b
155234	Dusp1	22464096	expression, and may play a role in the development of Th1 type immune response and anti-microbial defence. Dusp1 modulates the phosphorylation status of
155234	Dusp1	26019272	mRNA-destabilizing protein Zfp36 to regulate macrophage immune response to lipopolysaccharide. Bacterial cell wall glucosaminyl-muramyl dipeptides
189999	Cxcr4	26026270	(GMDPs) bind to the transcription factor Ybx1 to upregulate Nfkb2 and Cxcr4 gene expression. Bacterial cell wall glucosaminyl-muramyl dipeptides
172544	Nfkb2	26026270	(GMDPs) bind to the transcription factor Ybx1 to upregulate Nfkb2 and Cxcr4 gene expression. Ybx1 controls intracellular II6 mRNA levels in a cell type-specific manner, leading to functions that are
182925	Ybx1	25398005	dependent on the extracellular and intracellular distribution of Ybx1. Bacterial cell wall glucosaminyl-muramyl dipeptides (CMDPa) bind to the transmission factor Yby1 to
182925	Ybx1	26026270	(GMDPs) bind to the transcription factor Ybx1 to upregulate Nfkb2 and Cxcr4 gene expression.

			Antibody-dependent enhancement (ADE) of Dengue virus serotype 2 (DENV-2) elevates mature IL1B
75571	SYK	26032420	secretion via SYK signalling pathway in primary monocytes.
			Il7 is produced by intestinal epithelial cells in response to Citrobacter rodentium infection and plays a critical role in the protective immunity against this intestinal
130583	II7	26034215	attaching and effacing bacterium. Tlr5 forms a dimer upon binding to bacterial flagellin.
259066	Tlr5	22434932	(Demonstrated in human)
259066	Tlr5	22545147	Tlr5 is highly expressed in mucosal dendritic cells and Tlr5 signalling restricts regulatory T cell generation. Tlr5 deficiency in mice leads to a transient inability to
259066	Tlr5	22863420	manage proteobacteria which promotes chronic gut inflammation. Flagellin-specific IgG1 antibody response is induced
259066	Tlr5	24442437	through a Tlr5-, inflammasome-, and Myd88- independent pathway.
239000	1115	24442437	Flagellin induces Tlr5-dependent Il22 production and Nlrc4-dependent Il18 production to promote a protective gene expression program in intestinal
259066	Tlr5	25395539	epithelial cells and elimination of rotavirus-infected cells.
			Indirect Tlr5-dependent stimulation of airway conventional dendritic cells is essential to flagellin's
259066	Tlr5	26003491	mucosal adjuvant activity. IRF7 increases the expression of a broad range of IFN- stimulated genes including immunomodulatory
17225	IRF7	19152337	cytokines and genes involved in antigen processing and presentation. IRF7 is activated in response to virus infection and
17225	IRF7	15664995	stimulates the transcription of a set of cellular genes involved in host antiviral defence.
			IRF7 forms a complex with MYD88 and TRAF6 and this complex formation, as well as TRAF6-dependent
17225	IRF7	15361868	IRF7 ubiquitination, is required for TLR-mediated interferon (IFN)-alpha induction.
			IRF7 is involved in the innate immune recognition of Plasmodium falciparum AT-rich DNA and in the subsequent induction of type I IFNs. Mice lacking
17225	IRF7	21820332	Irf3/Irf7 are resistant to otherwise lethal cerebral malaria. (Demonstrated in mouse) Coronavirus engages papain-like proteases to escape
17225	IRF7	25505178	from the innate antiviral response of the host by inhibiting TP53-IRF7-IFNB1 signalling.
			Paramyxoviruses trigger the DNA-damage response, a pathway required for RPS6KA5 activation of phospho Ser 276 RELA formation to trigger the IRF7-DDX58
17225	IRF7	25520509	amplification loop necessary for mucosal interferon production.
17225	IRF7	25911105	AIP is a novel inhibitor of IRF7 and a negative regulator of innate antiviral signalling.

60515	AIP	25911105	AIP is a novel inhibitor of IRF7 and a negative regulator of innate antiviral signalling.
172144	Ager	22386596	Ager is a native receptor for complement component C1qa. (Demonstrated in human)
	0		S100a8/S100a9 are proinflammatory proteins that
172144	Ager	25911757	activate natural killer cells via Ager signalling. S100a9 forms a complex with S100a8 and the complex is the site of interplay between extracellular Ca(2+) entry and intra-phagosomal reactive oxygen species production. S100a8 :: S100a9 acts as Ca(2+) sensor in phagosomal ROS production.
168119	S100a9	21239714	
168119	S100a9	21382888	<ul> <li>S100a9-deficient murine neutrophils exhibited a reduce secretion of cytokines in response to Tlr4 stimulation.</li> <li>In contrast, S100a9-deficient dendritic cells showed an exacerbated release of cytokines after TLR stimulation.</li> <li>S100a9 has no effect on the inflammatory status of macrophages.</li> <li>S100a9 is strongly upregulated in neutrophils upon bacterial infection, and sequesters zinc as a mechanism of nutritional immunity. Salmonella typhimurium</li> </ul>
			overcomes this defence mechanism by expressing a
168119	S100a9	22423963	high affinity zinc transporter. S100A9 forms a heterodimer with S100A8 and is a key
168119	S100a9	23133376	player in protective innate immunity during Klebsiella pneumonia infection.
168119	S100a9	25911757	S100a8/S100a9 are proinflammatory proteins that activate natural killer cells via Ager signalling. S100a8 forms a complex with S100a9 and the complex is the site of interplay between extracellular Ca(2+) entry and intra-phagosomal reactive oxygen species production. S100A8 :: S100A9 acts as Ca(2+) sensor in phagosomal ROS production.
168077	S100a8	21239714	
			S100a8 is strongly upregulated in neutrophils upon bacterial infection, and sequesters zinc as a mechanism of nutritional immunity. Salmonella typhimurium overcomes this defence mechanism by expressing a
168077	S100a8	22423963	high affinity zinc transporter. S100A8 forms a heterodimer with S100A9 and is a key
168077	S100a8	23133376	player in protective innate immunity during Klebsiella pneumonia infection.
168077	S100a8	25911757	S100a8/S100a9 are proinflammatory proteins that activate natural killer cells via Ager signalling. Lcp2 is a critical determinant of natural killer (NK)-cell
161461	Lcp2	25929249	development and NK cell mediated elimination of missing-self target cells.
145267	Ticam2	21494017	Ticam2 deficiency results in the impairment of LPS- stimulated TNF-alpha protein translation.
145267	Ticam2	25505250	Adaptor proteins Ticam1 and Ticam2 have a novel function in Tlr2-mediated signal transduction.

			Ticam2 mediates antibacterial defence during Gram- negative pneumonia by inducing Ifng at the primary
145267	Ticam2	26065469	site of infection. Intracellular Sef/IL-17R (SEFIR) domain of Il17rd targets TIR adaptor proteins Myd88, Tirap, Ticam1,
145267	Ticam2	25808990	Ticam2 and Traf6 to inhibit TLR downstream signalling. Tirap Ser180Leu polymorphism is significantly associated with Behcet's disease in UK, but not Middle Eastern, patients. It is suggested that the Ser180Leu
			functional variant of Tirap will lead to greater cytokine production and tissue damage with persistence of mucosal lesions upon encounter with pathogens.
148134	Tirap	21705416	(Demonstrated in human) Intracellular Sef/IL-17R (SEFIR) domain of Il17rd targets TIR adaptor proteins Myd88, Tirap, Ticam1,
148134	Tirap	25808990	Ticam2 and Traf6 to inhibit TLR downstream signalling. Intracellular Sef/IL-17R (SEFIR) domain of Il17rd targets TIR adaptor proteins Myd88, Tirap, Ticam1,
140856	Il17rd	25808990	Ticam2 and Traf6 to inhibit TLR downstream signalling. RNA cleavage products, catalyzed by RNASEL, bind
21550	DHX33	25816776	to DHX33 to facilitate the formation of a complex with MAVS and NLRP3 during viral infection. RNASEL, antiviral endoribonuclease, is the terminal
			component of an RNA decay pathway that is an important mediator of IFN-induced antiviral activity and is required for the optimal induction of pro-
105248	RNASEL	19075243	inflammatory cytokines that play essential roles in host defence from bacterial pathogens. RNASEL regulates the expression of the endolysosomal protease, cathepsin-E, and endosome-
			associated activities, that function to eliminate internalized bacteria and may contribute to RNASEL
105248	RNASEL	19075243	antimicrobial action. RNASEL-mediated cleavage of Hepatitis C virus (HCV) RNA generates suppressor of virus RNA (avRNA) that activates DDX58 (BIC I) thus
105248	RNASEL	20833746	(svRNA) that activates DDX58 (RIG-I), thus propagating innate immune signalling to the interferon (IFN)-beta gene. RNASEL cleaves RNA during viral infections and the
			cleavage products induces the RIG-I pathway and production of IFNB gene. In addition, RNASEL is implicated in the protection of central nervous system against viral-induced demyelination. A broader role in innate immunity is suggested by involvement of RNASEL in cytokine induction and endosomal
105248	RNASEL	21190483	pathways that suppress bacterial infections.

RNA cleavage products, catalyzed by RNASEL, bind to DHX33 to facilitate the formation of a complex with MAVS and NLRP3 during viral infection.

105248	RNASEL	25816776	
11512	IL32	25820174	IL32 enhances host immunity to Mycobacterium tuberculosis.
24117	ELAVL1	14981256	ELAVL1 is a protein that binds to specific mRNA subsets, preferentially within 3' untranslated regions, and is a pivotal posttranscriptional regulator of gene expression.
			ELAVL1 is an RNA-binding protein can stabilize and/or regulate the translation of target mRNAs, thereby affecting the cellular responses to immune, proliferative, and damaging agents. It has a broad anti- apoptotic function where it increases the stability of a target mRNA encoding the pro-survival deacetylase SIRT1 and promotes the expression of mRNAs encoding BCL2 and MCL1, two major anti-apoptotic
24117	ELAVL1	17534146	effectors. ELAVL1 is required for the stabilization of IFNB1 mRNA, and suppression of ELAVL1 leads to impaired expression of IFNB1 in response to poly(I:C)
24117	ELAVL1	25678110	treatment. CYLD is a negative regulator of DDX58(RIG-I)- mediated antiviral response by removing Lys 63-linked
30697	CYLD	18636086	polyubiquitin chains from RIG-I and TBK1. CYLD is a crucial negative regulator of innate immune
30697	CYLD	18643924	response in Escherichia coli pneumonia. CYLD is a deubiquitinase that counteracts E3 ligases and therefore play a prominent role in the downregulation of NF-ΰB signalling and homeostasis.
30697	CYLD	21119682	
30697	CYLD	21498625	CYLD is a deubiquitinase that act as a negative regulator of TLR3 induction in response to LPS. CYLD plays a key role in Type I IFN receptor signalling during vesicular stomatitis virus (VSV) infection. In the absence of CYLD, IFN-beta is ineffective in the induction of antiviral genes.
30697	CYLD	21946435	(Demonstrated in mice) The E3 ligase ITCH and deubiquitinase CYLD act
30697	CYLD	22057290	together to regulate TAK1 and inflammation. MIR362 promotes natural killer-cell function by the
30697	CYLD	25909817	down-regulation of CYLD. MIR362 promotes natural killer-cell function by the
126785	MIR362	25909817	down-regulation of CYLD. Mir125a regulates the innate host defense by inhibiting the activation of autophagy and antimicrobial effects against Mycobacterium tuberculosis through targeting
200543	Uvrag	25917095	Uvrag.

			Mir125a regulates the innate host defense by inhibiting the activation of autophagy and antimicrobial effects
224267	Mir125a	25917095	against Mycobacterium tuberculosis through targeting Uvrag. Bordetella pertussis CyaA toxin plays a role in evading
187635	Ptpn6	25876760	nitric oxide-mediated killing in macrophages through a cAMP-dependent activation of the Ptpn6 phosphatase. CTSK is a cathepsins, which are key modulators of cell
102192	CTSK	18762176	death and inflammatory responses. Cleavage of CAMP by cathepsins CTSS and CTSK impairs its antimicrobial activity against Pseudomonas
102192	CTSK	25884905	aeruginosa and Staphylococcus aureus. CTSS is a member of the Cathepsins protein family,
102189	CTSS	18762176	<ul> <li>which are key modulators of cell death and inflammatory responses.</li> <li>CTSS is an endosomal and lysosomal protease that is upregulated during various inflammatory disorders.</li> <li>TLR2, 3, 4 ligand engagement increases the proteolytic activities of CTSS in macrophages. (Demonstrated in murine model)</li> </ul>
102189	CTSS	21145045	Cleavage of CAMP by cathepsins CTSS and CTSK
102189	CTSS	25884905	impairs its antimicrobial activity against Pseudomonas aeruginosa and Staphylococcus aureus. Mir328 is a key element of the host response to pulmonary infection with non-typeable Haemophilus influenzae and its inhibition in macrophages augments
223547	Mir328	25894560	phagocytosis, the production of reactive oxygen species, and microbicidal activity. MIR328 is a key element of the host response to pulmonary infection with non-typeable Haemophilus influenzae and its inhibition in macrophages augments
126687	MIR328	25894560	phagocytosis, the production of reactive oxygen species, and microbicidal activity. CAMP (LL-37), at sufficiently low concentrations, is able to reduce fungal infectivity by inhibiting C. albicans adhesion to plastic surfaces, oral epidermoid cells, and the urinary bladders of female mice. The inhibitory effects of LL-37 on cell adhesion and aggregation were mediated by its preferential binding
201591	Camp	21448240	to mannan and chitin in the fungal cell wall. (Demonstrated in human) Camp (LL-37) translocates across the E. coli outer
201591	Camp	21464330	membrane and halts bacterial growth by interfering cell wall biogenesis. (Demonstrated in human) Camp protects against colitis induction in mice. The increased expression of Camp in monocytes involves
201591	Camp	21762664	the activation of Tlr9/ERK signalling pathway by bacterial DNA. Camp expression is induced upon endoplasmic
201591	Camp	21832078	reticulum stress via NF-kB-C/EBP-alpha activation. (Demonstrated in human)

201591	Camp	22031815	mCRAMP (Camp) reduces influenza A viral load and disease severity in mice. The transcription factor Zfp423 is necessary for
201591	Camp	25554785	adipocyte activation and impaired adipogenesis is observed in Zfp423(nur12) mice. Camp plays an important role in the innate immune response against pathogens in bacterial central nervous
201591	Camp	25896094	<ul> <li>response against pathogens in bacterial central introductions</li> <li>system infections.</li> <li>Exogenous II4 was sufficient to drive the accumulation of tissue macrophages through self-renewal revealing that the expansion of innate cells necessary for pathogen control or wound repair can occur without recruitment of potentially tissue-destructive inflammatory cells.</li> </ul>
172024	Il4	21566158	IL4 attenuates Th1-chemokines expression at the site
172024	Il4	23991011	of inflammation reducing Th1 lymphocyte recruitment and limits pathogen clearance Nlrp12 is an intrinsic negative regulator of T-cell- mediated immunity. Altered NF-kB regulation and Il4 production are key mediators of Nlrp12-associated
172024	I14	25888258	disease. Nlrp12 is an intrinsic negative regulator of T-cell- mediated immunity. Altered NF-kB regulation and Il4 production are key mediators of Nlrp12-associated
264083	Nlrp12	25888258	disease. DEFB103B inhibits cell wall biosynthesis in staphylococci by inhibiting those enzymes which use the bactoprenol bound cell wall building block Lipid II
6693	DEFB103B	20385753	as substrate. DEFB103B exerts anti-inflammatory activities by
6693	DEFB103B	21809339	specifically targeting TLR signalling pathways to transcriptionally repress pro-inflammatory genes. DEFB103B (hBD-3) is sequestered by extracellular DNA in Haemophilus influenzae biofilms to reduce its
6693	DEFB103B	22922323	antimicrobial activity. DEFB103B (hBD3) induces dendritic cell activation,
6693	DEFB103B	22951718	migration and polarization in the skin. DEFB103B (hBD3) is induced in leprosy type 1
6693	DEFB103B	23133681	reactions in keratinocytes. Exposure to ambient air pollution particulate matter deregulates the ability of the human type II alveolar epithelial cells (A549) to express the antimicrobial peptides HBD-2 (DEFB4A/DEFB4B) and HBD-3 (DEFB103A/DEFB103B) upon infection with Mycobacterium tuberculosis and increases intracellular
6693	DEFB103B	25847963	M. tuberculosis growth. DEFB103A inhibits cell wall biosynthesis in staphylococci by inhibiting those enzymes which use the bactoprenol bound cell wall building block Lipid II
6300	DEFB103A	20385753	as substrate.

DEFB103A is a beta-defensin with direct antimicrobial properties that contribute to local innate immune responses and it aids in combating microbial invasion by being chemotactic for a broad spectrum of leukocytes in a CCR6- and CCR2-dependent manner.

6300 DEFB103A 20483750 DEFB103A expression is inhibited in human epidermal keratinocytes under high glucose conditions, which in turn contributed to the frequent occurrences of infection associated with diabetic wounds.

> DEFB103A (hBD-3) is sequestered by extracellular DNA in Haemophilus influenzae biofilms to reduce its antimicrobial activity.

DEFB103A (hBD3) induces dendritic cell activation, migration and polarization in the skin.

6300DEFB103A(hBD3) is induced in leprosy type 1reactions in keratinocytes.

6300

6300

6283

6711

DEFB103A

DEFB4B

DEFB4

20483750

DEFB103A and RNASE7 are induced in human umbilical endothelial cells (HUVECs) by classical 6300 25637949 inflammatory cytokines such as: IFNG, IL1B and TNF. DEFB103A Exposure to ambient air pollution particulate matter deregulates the ability of the human type II alveolar epithelial cells (A549) to express the antimicrobial peptides HBD-2 (DEFB4A/DEFB4B) and HBD-3 (DEFB103A/DEFB103B) upon infection with Mycobacterium tuberculosis and increases intracellular 6300 DEFB103A 25847963 M. tuberculosis growth.

Exposure to ambient air pollution particulate matter deregulates the ability of the human type II alveolar epithelial cells (A549) to express the antimicrobial peptides HBD-2 (DEFB4A/DEFB4B) and HBD-3 (DEFB103A/DEFB103B) upon infection with Mycobacterium tuberculosis and increases intracellular 25847963 M. tuberculosis growth.

> DEFB4 is a beta-defensin with direct antimicrobial properties that contribute to local innate immune responses and it aids in combating microbial invasion by being chemotactic for a broad spectrum of leukocytes in a CCR6- and CCR2-dependent manner.

DEFB4A is upregulated in hypoxic<br/>microenvironments, which is characteristic of infected<br/>tissue.6711DEFB422427634EEEE to be the second second

6711DEFB422500651DEFB4A induction in keratinocytes is inhibited by<br/>Pseudomonas aeruginosa rhamnolipids.

6711DEFB425510212In alveolar epithelium, IL22 upregulates DEFB4A gene<br/>expression via STAT3.

6711	DEFB4	25847963	Exposure to ambient air pollution particulate matter deregulates the ability of the human type II alveolar epithelial cells (A549) to express the antimicrobial peptides HBD-2 (DEFB4A/DEFB4B) and HBD-3 (DEFB103A/DEFB103B) upon infection with Mycobacterium tuberculosis and increases intracellular M. tuberculosis growth. Mapk14 activation is blocked by Bacillus anthracis, resulting in the opening of a connexin ATP release channel and induction of macrophage death.
158983	Mapk14	21683629	Constitutive activation of Mapk14 interferes with inflammasome activation and II1b production, which compromises antimicrobial immunity. The Mapk14 pathway is an important contributor to
158983	Mapk14	21733175	microglial production of proinflammatory cytokines induced by LPS or beta-amyloid. Mapk14 mediates cytoskeletal remodelling and early
158983	Mapk14	22028692	<ul><li>spreading of lipopolysaccharide (LPS)-stimulated macrophages.</li><li>Mapk1 (ERK) and Mapk14 (p38) control the dynamic</li></ul>
158983	Mapk14	22447027	balance regulating neutrophil migration. (Demonstrated in human) Lysophosphatidic acid plays an anti-inflammatory role in macrophages by diminishing lipopolysaccharide-
158983	Mapk14	25783839	induced phosphorylation of Mapk14 and Akt1, as well as Rela nuclear translocation.
158983	Mapk14	25848864	Treml4 is an essential positive regulator of Tlr7 signalling. Treml4(-/-) macrophages are hyporesponsive to Tlr7 agonists and fail to produce type I interferons due to impaired phosphorylation of Stat1 by Mapk14 and decreased recruitment of Myd88 to Tlr7. Treml4 is an essential positive regulator of Tlr7 signalling. Treml4(-/-) macrophages are hyporesponsive to Tlr7 agonists and fail to produce type I interferons due to impaired phosphorylation of
189814	Trem14	25848864	Stat1 by Mapk14 and decreased recruitment of Myd88 to Tlr7.
44230	TLR7	18071655	<ul><li>TLR7 recognizes long single-stranded RNA and short double-stranded RNA.</li><li>TLR7 recognizes the single-stranded RNA viruses, vesicular stomatitis virus and influenza virus, resulting</li></ul>
44230	TLR7	15034168	in activation of co-stimulatory molecules and production of cytokines. TLR7-dependent production of inflammatory cytokines
44230	TLR7	14976261	can be induced by single-stranded RNA (ssRNA) molecules of non-viral origin. TLR7, 8 and 9 form a functional subgroup within the TLR family that recognizes pathogen-associated
44230	TLR7	14579267	molecular patterns in endosomal/lysosomal compartments.

44230	TLR7	20034855	TLR7/9-mediated innate immune responses via selected TLR pathways can be negatively regulated by a human microsatellite DNA-mimicking oligodeoxynucleotide with CCT repeats.
44230	ILK/	20034833	TLR7-dependent innate immune response is induced by free human T-cell leukemia virus 1 (HTLV-1) in killer plasmacytoid dendritic cells, resulting in high
44230	TLR7	20007807	production of IFN-alpha and TRAIL relocalization. TLR7 is expressed in C-fiber primary sensory neurons and is important for inducing itch (pruritus), but is not necessary for eliciting mechanical, thermal, inflammatory and neuropathic pain. (Demonstrated in murine model)
44230	TLR7	21037581	
			TLR7 signalling pathway plays a pivotal role in fungal pathogen recognition and is essential for the subsequent IFNB signalling. (Demonstrated in murine model)
44230	TLR7	21282509	,
			TLR7 requires proteolytic processing in endolysosome by asparagine endopeptidase and cathepsin in the endolysosome to initiate signalling. (Demonstrated in
44230	TLR7	21402738	murine model) TLR7 agonists, such as imidazoquinolines, accumulate in the MHC class II loading compartment - this pH- dependent localization is required for the activation of
44230	TLR7	21487111	plasmacytoid dendritic cells. TLR7 inflammatory signalling leads to cardiac fibrosis
44230	TLR7	21730058	in autoimmune associated congenital heart block. TLR7 and TLR8 are translocated from the endoplasmic reticulum to the endosome in the presence of antiphospholipid antibodies, as a consequence, plasmacytoid dendritic cells become dramatically sensitized to TLR7/8 agonists and this may play a role
44230	TLR7	21734241	in systemic autoimmunity. TLR7 is responsible for the detection of retroviruses and serves as a key checkpoint controlling the
44230	TLR7	21998589	development of germinal center B cells. (Demonstrated in mice)
44230	TLR7	22396599	TLR7 signalling induces autophagy in HIV-infected plasmacytoid dendritic cells; this process is necessary for the induction of IFN-alpha. Aberrant TLR7 activation induces Epstein-Barr viral
44230	TLR7	22952664	protein LMP1 expression, which exacerbates IFN production in lupus patients. IFN- $\hat{I} \gg 1$ is able to augment TLR-mediated B cell
44230	TLR7	26130701	activation, partially attributed to an upregulation of TLR7 expression SPHK1 is involved in toll-mediated human beta- defensin 2 (HBD-2) regulation in oral keratinocytes,
69650	SPHK1	20634980	which also involves the activation of PI3K, AKT, GSK3B (GSK-3beta) and ERK 1/2.

69650	SPHK1	20661259	SPHK1 activation, mediated by TLR4, is found to be critical for the redox-dependent activation of HIF- 1alpha and ASK1, as well as for the prevention of LPS- induced activation of caspase 3 and the expression of pro-inflammatory cytokine interleukin-6. (S)-methyl 2-(hexanamide)-3-(4-hydroxyphenyl)
69650	SPHK1	26113114	propanoate (MHP) activates SPHK1, which also stimulates CAMP production and enhances epidermal antimicrobial defence. Peli1 is an innate immune regulator in the central
153059	Peli1	26131354	nervous system that modulates the threshold of Type I interferon responses against viral infections. FCN3, as well as ficolins FCN1 and FCN2, in serum are associated with MBL-associated serine protease (MASP) to form a complex and this complex binds to carbohydrates present on the surface of a variety of Gram-positive and Gram-negative bacteria through
94882	FCN3	20375620	ficolin, initiating complement activation via the lectin pathway. H-Ficolin (FCN3), an innate immune opsonin, participates in Aspergillus fumigatus defence through the activation of the lectin complement pathway,
94882	FCN3	26133042	enhancement of fungus-host interactions and modulation of immune responses. Group 2 innate lymphoid cell (ILC2)-intrinsic II33 signalling and Icosl expression promote regulatory T cell accumulation, whereas the inflammatory cytokine
168471	Icosl	26092469	Ifng counter-regulated the effects of II33, in part through direct effects on ILC2s. The membrane-modulating enzyme SMPDL3B is a negative regulator of TLR signalling that functions at
194305	Smpdl3b	26095358	the interface of membrane biology and innate immunity. Glycogen synthase kinase $3/Ctnnb1$ ( $\hat{1}^2$ -catenin) axis is
205243	Ctnnb1	26100021	required for optimal induction of antiviral innate immunity. Glycogen synthase kinase 3/Ctnnb1 (Î <sup>2</sup> -catenin) axis is
156635	Gsk3a	26100021	required for optimal induction of antiviral innate immunity. Gsk3b is a regulator of LPS-mediated septic shock. Gsk3b deficiency results in the attenuation of endotoxemia.
159859	Gsk3b	21515258	Gsk3b functions downstream of Tlr2-stimulation to
159859	Gsk3b	22218715	induce the expression of the monocyte chemoattractant protein 1, Ccl2. Glycogen synthase kinase 3/Ctnnb1 (Î <sup>2</sup> -catenin) axis is
159859	Gsk3b	26100021	required for optimal induction of antiviral innate immunity. CTNNB1 binds to NFKB1 and regulates expression of
27347	CTNNB1	17704137	C-Reactive protein (CRP) after TNF (TNF-alpha) treatment.

			CTNNB1 interacts with LRRFIP1, promoting the activation of CTNNB1, which increases IFN-beta expression by binding to the C-terminal domain of the transcription factor IRF3 and recruiting the
27347	CTNNB1	20453844	acetyltransferase EP300 to the IFN-beta enhanceosome via IRF3.
27347	CTNNB1	23785285	Stabilization of CTNNB1 upon virus infection negatively regulates antiviral innate immunity. Glycogen synthase kinase 3/CTNNB1 (Î <sup>2</sup> -catenin) axis is required for optimal induction of antiviral innate
27347	CTNNB1	26100021	immunity. GSK3B is a cytoplasmic serine/threonine protein
51663	GSK3B	17912008	kinase that regulates NF-kappaB activation and the proliferation and survival of pancreatic cancer cells. GSK3B inhibits MEKK4 activity and prevents its activation of JNK and p38, thus controlling MEKK4
51663	GSK3B	17726008	dimerization both positively and negatively by regulating its interaction with specific proteins. GSK3B activation is accelerated by TLR4 which leads to deterioration of serum-deprivation-induced apoptosis and beta-arrestin 2 represents an inhibitory effect on the TLR4-mediated apoptotic cascade, through controlling
51663	GSK3B	20497256	the homeostasis of activation and inactivation of GSK3B. GSK3B is a regulator of LPS-mediated septic shock. GSK3B deficiency results in the attenuation of endotoxemia. (Demonstrated in murine model)
51663	GSK3B	21515258	GSK3B functions downstream of TLR2-stimulation to
51663	GSK3B	22218715	induce the expression of the monocyte chemoattractant protein 1, CCL2. (Demonstrated in mice) Glycogen synthase kinase 3/CTNNB1 (Î <sup>2</sup> -catenin) axis
51663	GSK3B	26100021	is required for optimal induction of antiviral innate immunity. Glycogen synthase kinase 3/CTNNB1 (Î <sup>2</sup> -catenin) axis
53956	GSK3A	26100021	is required for optimal induction of antiviral innate immunity. TLR8 expression and function is highly up-regulated in colonic epithelium from patients with active inflammatory bowel disease (IBD), suggesting that TLR8 aignelling is important in the patheoremic of
44288	TLR8	18985539	TLR8 signalling is important in the pathogenesis of IBD. Toll-like receptor 8 (TLR8), 7 and 9 form a functional subgroup within the TLR family that recognizes
44288	TLR8	14579267	pathogen-associated molecular patterns in endosomal/lysosomal compartments. TLR8 and TLR7 bind single-stranded RNA, stimulating dendritic cells (DCs) and macrophages to
44288	TLR8	14976262	secrete interferon-alpha and pro-inflammatory, as well as regulatory, cytokines.

44288	TLR8	17264163	TLR8 signalling strongly promotes inflammatory lipid mediator biosynthesis, providing novel insights on innate immune response to viral infections. TLR8 is activated in human monocytic cells following Helicobacter pylori phagocytosis and TLR8 single nucleotide polymorphism play a role in the modulation
44288	TLR8	20652908	of TLR8-dependent microbicidal response of infected macrophages. TLR7 and TLR8 are translocated from the endoplasmic reticulum to the endosome in the presence of antiphospholipid antibodies, as a consequence, plasmacytoid dendritic cells become dramatically
44288	TLR8	21734241	sensitized to TLR7/8 agonists and this may play a role in systemic autoimmunity. TLR8 binding of HIV ssRNA induces endosomal acidification and chromatin remodeling at the TNF- alpha promoter to promote TNF-alpha release in
44288	TLR8	22393042	infected macrophages. TLR8 binds to exosomal MIR21 and MIR29A secreted by tumour cells and initiates a prometastatic
44288	TLR8	22753494	inflammatory response. TLR8 plays a pathogenic role in disease whereby its expression is increased in patients with systemic
44288	TLR8	24277153	arthritis and is correlated with the elevation of IL1B levels and disease status. TLR8 binds degradation products of single-stranded RNA at two distinct sites. One site prefers uridine mononucleosides and the other site prefers short
44288	TLR8	25599397	oligonucleotides. TLR8-dependent detection of bacterial RNA is critical
44288	TLR8	26101323	for triggering monocyte activation in response to infection with Streptococcus pyogenes. TRIM5 requires UBE2W, UBE2N and UBE2V2
21151	UBE2V2	26101372	enzymatic activities to inhibit retroviral DNA synthesis UBE2N abalation resulted in defective B cell development and in impaired B cell and macrophage
51220	UBE2N	16862162	activation. TRIM5 requires UBE2W, UBE2N and UBE2V2
51220	UBE2N	26101372	enzymatic activities to inhibit retroviral DNA synthesis TRIM5 requires UBE2W, UBE2N and UBE2V2
25727	UBE2W	26101372	enzymatic activities to inhibit retroviral DNA synthesis TRIM5 is upregulated by Type I and Type II interferons and have been found to restrict viral
27690	TRIM5	21131187	replication by modulating the RIG-I pathway. TRIM5 is an innate intracellular HIV restriction factor
27690	TRIM5	21734563	that is upregulated by type I interferons. TRIM5 was identified in a systematic screen for
27690	TRIM5	23438823	positive regulators of innate immune responses. TRIM5 requires UBE2W, UBE2N and UBE2V2
27690	TRIM5	26101372	enzymatic activities to inhibit retroviral DNA synthesis

			OAS3 is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the OAS3 activates RNASEL to cleave ssRNA. The OAS/RNASEL pathway triggers the RIG-I pathway and induce IFNB production.
58353	OAS3	21190483	Antiviral activity of the $2\hat{a} \in 2$ , $5\hat{a} \in 2$ -oligoadenylate
58353	OAS3	26063222	synthetase (OAS) gene variants against dengue virus shows strong and serotype-specific disparities. OAS1 is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the OAS1 activates RNASEL to cleave ssRNA. The OAS/RNASEL pathway triggers the RIG-I pathway and induce IFNB production.
58231	OAS1	21190483	F
58231	OAS1	23319625	OAS1 is a cytoplasmic dsRNA sensor. Antiviral activity of the $2 \hat{a} \in 2$ , $5 \hat{a} \in 2$ -oligoadenylate synthetase (OAS) gene variants against dengue virus
58231	OAS1	26063222	shows strong and serotype-specific disparities.
210514	Pycard	21439959	Pycard is a component of the inflammasome and is required for inflammation in acute pancreatitis. Nlrp3/Pycard inflammasome activation following human respiratory syncytial virus infection is dependent on the activation of Tlr2/Myd88/NF-kB and
210514	Pycard	22295065	reactive oxygen species/potassium efflux. (Demonstrated in human)
210514	Pycard	23302887	<ul><li>PYCARD is an essential regulator of inflammatory responses in West Nile virus encephalitis.</li><li>Phosphorylation of the inflammasome adaptor Pycard (ASC) controls inflammasome activity through the formation of ASC specks. The NLRP3 and AIM2 inflammasomes require Syk and Mapk8 (JNK) for their</li></ul>
210514	Pycard	24185614	full activity . Activation of the Nlrp3 inflammasome is detrimental during leishmaniasis. Mice lacking the inflammasome components Nlrp3, Pycard, Casp1 exhibit defective Il1b and Il18 production at the infection site and are
210514	Pycard	25689249	resistant to cutaneous Leishmania major infection. Mice lacking Pycard display attenuated pulmonary hypertension and right ventricle remodelling in response to hypoxia and this is accompanied by
210514	Pycard	26071556	blunted inflammasome activation. Psen2 deficiency is paralleled by reduced transcription of Tlr4 mRNA and loss of LPS-induced Tlr4 mRNA
207109	Psen2	26081153	transcription regulation. Stmn1 overexpression impacts microtubule stability, impairs cell spreading, reduces activation-associated phenotypes and reduces complement receptor 3 (CR3)-
196306	Stmn1	26082487	mediated phagocytosis and cellular activation.

MAP3K7 (TAK1) is a member of the mitogenactivated protein kinase kinase kinase family that, together with its activator TAK1-binding protein 1 (TAB1), activates the IKK signallosome and thus regulates NF-kappaB activation.

10187861 MAP3K7 is an essential intermediate of NOD2 signalling where MAP3K7 deletion completely abolishes muramyl dipeptide (MDP)-NOD2 signalling, activation of NF-kappaB and MAPKs, and the subsequent induction of cytokines/chemokines in 17965022 keratinocytes.

MAP3K7 mediates the activation signal from TLRs to nuclear factor-kappaB in lipopolysaccharide-stimulated macrophages.

MAP3K7 acts as an upstream activating kinase for IKBKB (IKK-beta) and MAPK8 (JNK), but not CHUK (IKK-alpha), revealing a specific role of MAP3K7 in 16260493 inflammatory signalling pathways.

> MAP3K7 (TAK1) functions as a mediator in the signalling pathway of TGF-beta superfamily members.

MAP3K7 induces NF-kappaB activation through a 94374 MAP3K7 9480845 MAP3K14 (NIK)-independent signalling pathway.

> MAP3K7 links TRAF6 to the MAP3K14 (NIK)-IKK cascade in the IL-1 signalling pathway where activated MAP3K7 phosphorylates NIK, which stimulates IKK-10094049 alpha activity.

> MAP3K7 is a target for glucocorticoids that integrates their anti-inflammatory action in innate immunity 20065289 signalling pathways.

> > MAP3K7 plays a central role in controlling nuclear and cytoplasmic signalling cascades in primary neutrophils where it constitutively associates with the IKBKB (Ikappa-B kinase) complex in the nucleus and cytoplasm, impacting downstream signalling processes.

MAP3K7 activation is impaired during endotoxin tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of anti-inflammatory or antimicrobial mediators.

MAP3K7 polyubiquitination is essential for the activation of NF-kB signalling downstream of TNF receptor, IL1 receptor and TLR4.

MAP3K7 is necessary for the neutrophil priming effect of leukotriene B (4) to enhance TLR stimulation. 22843747 (Demonstrated in mice)

> MAP3K7 (TAK1) Ser412 phosphorylation is regulated by PRKACA and PRKX, and is essential for proper signalling, as well as proinflammatory cytokine induction by TLR/IL-1R activation.

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## **MAP3K7**

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94374	MAP3K7	25371197	ECSIT binds to MAP3K7 and TRAF6 to form a complex that plays a pivotal role in activating TLR4-mediated NF-kB signalling. Endotoxin tolerance re-programs TLR4 signalling via suppression of PELI1, a positive regulator of MyD88-
94374	MAP3K7	26082489	and TIR domain-containing adapter inducing IFN- $\hat{1}^2$ (TRIF)-dependent signalling that promotes K63-linked polyubiquitination of IRAK1, TBK1, and TAK1. IRAK1 plays an essential role in Toll-like receptor/interleukin-1 receptor (TLR/IL-1R) - associated NF-kappaB activation through its
90782	IRAK1	18276832	involvement in IKK activation, which then leads to subsequent IkappaB degradation and NF-kappaB nuclear translocation and both cytosolic and nuclear actions of IRAK1 participate in the activation of NF- kappaB-dependent transcriptional events. IRAK1 and IRAK4 play key roles in a signalling pathway by which bacterial infection or interleukin-1 (IL1) trigger the production of inflammatory mediators. Pellino isoforms are the E3 ubiquitin ligases that mediate the IL1-stimulated formation of K63-pUb- IRAK1 in cells, which contribute to the activation of
90782	IRAK1	17997719	IKBKB and NF-kappaB, as well as other signalling pathways dependent on IRAK1 and IRAK4. IRAK1 is a component of a novel signal transduction
90782	IRAK1	10224059	pathway through which TNF receptor activates NF- kappaB-dependent gene expression. IRAK1 plays an essential proximal role in coordinating
90782	IRAK1	9625767	multiple IL1 signalling pathways for optimal induction of cellular responses.
90782	IRAK1	20044140	IRAK1 functionally associates with PKC-epsilon and VASP in the regulation of macrophage migration. IRAK1 and MYD88 autosomal recessive deficiencies
90782	IRAK1	21057262	impair Toll-like receptor (TLR)- and interleukin-1 receptor-mediated immunity and predispose patients to recurrent life-threatening bacterial diseases, such as invasive pneumococcal disease in particular, in infancy and early childhood, with weak signs of inflammation. IRAK1 is polyubiquitinated and disassembled during endotoxin tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of anti- inflammatory or anti-microbial mediators.
90782	IRAK1	21220427	
90782	IRAK1	22033459	IRAK1 mediates the proteasome-dependent degradation of TRAF6 and acts as a negative regulator of TLR-signalling. (Demonstrated in mice) IRAK1 accumulation triggers ischemia-induced
90782	IRAK1	23143987	inflammation in the small intestine. MIR146A is a potent negative regulator of the innate
90782	IRAK1	24670381	immune response in keratinocytes through downregulation of the IRAK1/TRAF6/NFI <sup>o</sup> B pathway.

			Endotoxin tolerance re-programs TLR4 signalling via suppression of PELI1, a positive regulator of MyD88- and TIR domain-containing adapter inducing IFN- $\hat{1}^2$
90782	IRAK1	26082489	(TRIF)-dependent signalling that promotes K63-linked polyubiquitination of IRAK1, TBK1, and TAK1. MYD88 is a Toll/IL-1R homology (TIR) domain containing adaptor which recruits IRAK1 possibly
25713	MYD88	11976320	through IRAK4. MYD88 can interact with bacterial TIR domain containing-proteins (Tcps) identified in Escherichia coli CFT073 (TcpC) and Brucella melitensis (TcpB) and interfere with MYD88-dependent pathway, thus
25713	MYD88	18327267	suppressing innate immunity and increasing virulence. MYD88 binding with interleukin-1 (IL-1) receptor (IL1R1) is required for inducing endocytosis of IL1R1
25713	MYD88	16354686	following ligand binding. MYD88 is a key adaptor/regulator molecule for the
25713	MYD88	9734363	Toll/IL-1R family of receptors for innate immunity. MYD88 interacts with the IL-1 receptor and blocks NF-kappaB activation induced by IL-1, but not by
25713	MYD88	9430229	TNF. TLR-2/MyD88/PI3K/Rac1/Akt pathway mediates LTA-induced MAPKs activation, which in turn initiates the activation of NF-kappaB, and ultimately
25713	MYD88	20167866	induces cPLA2/COX-2-dependent PGE2 and IL-6 generation. MYD88 plays a critical role in reverse cholesterol transport in vitro and in vivo, in part through promoting ATP-binding cassette A1 transporter upregulation, coupling cholesterol trafficking to inflammation through MYD88 and identifying innate
25713	MYD88	20519121	<ul> <li>immunity as a physiologic signal in cholesterol homeostasis.</li> <li>MYD88 and IRAK1 autosomal recessive deficiencies impair Toll-like receptor (TLR)- and interleukin-1 receptor-mediated immunity and predispose patients to recurrent life-threatening bacterial diseases, such as invasive pneumococcal disease in particular, in infancy</li> </ul>
25713	MYD88	21057262	and early childhood, with weak signs of inflammation. MYD88 is essential in restricting TLR3 signalling and the host protection from unwanted immunopathologies associated with excessive production of IFNB1. MYD88 inhibits TLR3 signalling by impairing IKBKE-mediated induction of IRF3, and consequently
25713	MYD88	21248248	<ul><li>the expression IFNB1 and CCL5.</li><li>MYD88 is activated by MHC class II in response to staphylococcal enterotoxins and is crucial for the</li></ul>
25713	MYD88	21283748	induction of pro-inflammatory cytokines.

			MYD88 is a key signalling adapter in TLR signalling. MYD88 aggregates in the cell as distinct foci and co- localizes with IRAK4 in these Myddosomes - the
25713	MYD88	21325272	formation of which is required for MYD88 function. MYD88 is required in dendritic cells stimulated with TLR9 ligand for the enhancement of T cell-dependent antibody response. In addition, MYD88 is required in B cells to facilitate strong anti-viral antibody
25713	MYD88	21353603	responses. (Demonstrated in murine model) MYD88 deficient macrophages displayed impaired interaction with fungal yeast cells and produced low levels of pro-inflammatory cytokines. MYD88 signalling is important in the activation of fungicidal mechanisms and the induction of protective innate immune responses against P. brasiliensis.
25713	MYD88	21422180	(Demonstrated in murine model) MYD88 mediates cytoskeletal remodelling and late spreading of lipopolysaccharide (LPS)-stimulated
25713	MYD88	22028692	macrophages. (Demonstrated in mice) MYD88-dependent recruitment of inflammatory monocytes and dendritic cells to the lungs are key initial cellular responses required for early protection from Burkholderia mallei infection. (Demonstrated in
25713	MYD88	22025508	mice) MYD88 deficiency results in delayed recruitment of phagocytes and defective production of
25713	MYD88	22386951	proinflammatory cytokines in response to Salmonella infection. (Demonstrated in mice) MYD88 signalling in intestinal epithelial cells is arresial for the maintenance of out microbioto
25713	MYD88	22491177	crucial for the maintenance of gut microbiota homeostasis. (Demonstrated in mice) MYD88 mediated production of reactive oxygen species (ROS) is essential for the induction of IL12 by
25713	MYD88	22536449	lactic acid bacteria. (Demonstrated in mice) Following NOD2 activation, IRF4 interacts with MYD88, TRAF6, and RIPK2 and downregulates K63-
25713	MYD88	24670424	linked polyubiquitinylation of RICK and TRAF6 leading to disruption of NFkB activation pathways. Endotoxin tolerance re-programs TLR4 signalling via suppression of PELI1, a positive regulator of MyD88- and TIR domain-containing adapter inducing IFN-1 <sup>2</sup> (TRIF)-dependent signalling that promotes K63-linked
25713	MYD88	26082489	polyubiquitination of IRAK1, TBK1, and TAK1. PELI1 is a ubiquitin ligase that facilitates TRIF- dependent Toll-like receptor signalling and pro-
53855	PELI1	19734906	inflammatory cytokine production. PELI1 (pellino) isoforms are E3 ubiquitin ligases that mediate the IL-1-stimulated formation of K63-pUb- IRAK1 in cells, contributing to the activation of
53855	PELI1	17997719	IKBKB and NF-kappaB, as well as other signalling pathways dependent on IRAK1 and IRAK4.

			Discrete regions of PELI1 are bound by SMAD6 and SMAD7 via their MH2 domains to mediate TGF-
53855	PELI1	20171181	beta1-induced negative regulation of IL-1R/TLR signalling. PELI1 is an adaptor protein involved in IL1R/TLR
53855	PELI1	21120624	signaling. PELI1 is sumoylated by SUMO1at 5 lysine residues, and binds to the SUMO-conjugating enzyme UBE2I.
			Endotoxin tolerance re-programs TLR4 signalling via suppression of PELI1, a positive regulator of MyD88- and TIR domain-containing adapter inducing IFN- $\hat{1}^2$
53855	PELI1	26082489	(TRIF)-dependent signalling that promotes K63-linked polyubiquitination of IRAK1, TBK1, and TAK1. Bacterial-derived monosaccharide heptose-1,7- bisphosphate (HBP) is a pathogen-associated molecular
34438	TIFA	26068852	pattern (PAMP) that activates TIFA-dependent immunity to Gram-negative bacteria The transcription factor Rora is critical for the
181478	Rora	22267218	development of nuocytes and the mounting of innate type 2 immunity against parasitic worms. Nlrp3 is a novel molecular target for melatonin which
181478	Rora	26045547	requires Rora to blunt the NFkB/ NLRP3 connection during sepsis. Irak1 is polyubiquitinated and disassembled during
			endotoxin tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of anti-
154747	Irak 1	21220427	inflammatory or anti-microbial mediators. Irak1 mediates the proteasome-dependent degradation
154747	Irak 1	22033459	of Traf6 and acts as a negative regulator of TLR- signalling.
154747	Irak 1	23143987	IRAK1 accumulation triggers ischemia-induced inflammation in the small intestine. Mir146 attenuates sepsis-induced cardiac dysfunction by preventing NF Î °B activation, inflammatory cell
154747	Irak 1	26048146	infiltration, and inflammatory cytokine production via targeting of Irak1 and Traf6 in both cardiomyocytes and inflammatory monocytic cells
134747	nuki	20040140	Human papillomaviruses impair the acetylation of NFΰB/RelA K310 in keratinocytes by augmenting the
36786	IFRD1	26055519	expression of interferon-related developmental regulator 1 (IFRD1) in an EGFR-dependent manner. EGFR is the receptor for epidermal growth factor (EGF) and signalling through growth factor receptors
16933	EGFR	11470832	controls diverse cell functions such as proliferation, migration, and differentiation. EGFR-induced cell migration is mediated
16933	EGFR	15284024	predominantly by the JAK-STAT pathway in primary esophageal keratinocytes.

1(022	ECED	10770126	EGFR and TLR4 are activated by neutrophil elastase (ELA2) to produce IL8 through a novel
16933	EGFR	18772136	metalloprotease pathway. Human papillomaviruses impair the acetylation of NFΰB/RelA K310 in keratinocytes by augmenting the
16933	EGFR	26055519	expression of interferon-related developmental regulator 1 (IFRD1) in an EGFR-dependent manner. PDE12 negatively regulates the innate immune response and inhibitors of PDE12 show increased IFN
			induced 2â€,5†- oligoadenylate and antiviral activities.
41300	PDE12	26055709	
408826	LYN	20385881	The LYN/PI3K module negatively regulates activation of murine macrophages while Inpp5d (SHIP-1) promotes it.
400020		20303001	Overexpression of LYN results in endotoxin
			hypersensitivity due to the increased activation of dendritic cells leading to an over-production of IFNG
408826	LYN	22491248	by natural killer cells. (Demonstrated in mice) LYN-dependent phosphorylation of the p110 catalytic subunit of PI 3-kinase is essential to the control of PI
			3-kinase biological activity upstream of AKT and
408826	LYN	26055819	thereby to the transactivation of NFΰB. MB21D1 (cGAS) is a cytosolic DNA sensor and is
92358	MB21D1	23707061	required for the induction of the interferon response. Cytosolic DNA sensor cyclic GMP-AMP synthase
92358	MB21D1	24284630	(cGAS, also known as MB21D1) is pivotal in protecting the host from both DNA and RNA viruses Cytosolic DNA sensor cGAS (MB21D1) is essential in
92358	MB21D1	24269171	human dentritic cells for innate sensing of HIV-1 and HIV-2.
			Cytosolic RNA:DNA hybrids are sensed by the MB21D1-TMEM173 (cGAS-STING) pathway of the
92358	MB21D1	25425575	innate immune system. IFI16 and MB21D1 interact and cooperate during
92358	MB21D1	25831530	herpes simplex virus infection to initiate innate signalling.
			PQBP1 directly binds to reverse-transcribed HIV-1 DNA and interacts with MB21D1 to initiate an IRF3-
92358	MB21D1	26046437	dependent innate response. PQBP1 directly binds to reverse-transcribed HIV-1
64349	PQBP1	26046437	DNA and interacts with MB21D1 to initiate an IRF3- dependent innate response. TNF (TNF-alpha) is an important mediator of
			inflammation, apoptosis, and the development of
300259	TNF	16199883	secondary lymphoid structures and LRRFIP1 represses TNF expression.

TNF pre-treated macrophages exhibit endotoxin tolerance, i.e. less cytokine production, towards LPS challenge. TNF-mediated cross-tolerization is mediated by suppression of LPS-induced signalling and chromatin remodelling.

300259	TNF	21602809	enfoliatin remoderning.
300259	TNF	21611132	TNF is essential to mount an acute inflammatory response to dsDNA in the endothelium. (Demonstrated in murine model) IFN gamma creates a primed chromatin environment in
300259	TNF	24012417	macrophages to augment TLR-induced TNF transcription Secreted CCNA2 (CCN1) promotes anti-inflammatory cytokine IL10 release from epithelial cells via integrin
300259	TNF	25005359	$\hat{I}\pm V\hat{I}^2$ 6-PKC, and this subsequently suppresses TNF, CXCL2 and neutrophil infiltration in the lungs. DEFB103A and RNASE7 are induced in human umbilical endothelial cells (HUVECs) by classical
300259	TNF	25637949	inflammatory cytokines such as: IFNG, IL1B and TNF. RNASE7 is produced by airway epithelial basal cells in response to cigarette smoke exposure.
2528	RNASE7	25712218	
2528	RNASE7	25637949	DEFB103A and RNASE7 are induced in human umbilical endothelial cells (HUVECs) by classical inflammatory cytokines such as: IFNG, IL1B and TNF. AIM2 recognizes cytosolic double stranded DNA and
103863	AIM2	19158675	forms a caspase-1-activating inflammasome with PYCARD (ASC). AIM2 has a critical role in host innate immunity to
103863	AIM2	20351693	intracellular pathogens where it is a crucial sensor of F. tularensis infection.
103863	AIM2	20351692	<ul><li>AIM2 inflammasome is essential for host defence against cytosolic bacteria and DNA viruses.</li><li>AIM2 is a newly discovered pattern recognition receptor (PRR) involved in the sensing of dangerous</li></ul>
103863	AIM2	20401524	cytosolic DNA produced by infection with DNA viruses. AIM2 is required for innate immune recognition of Francisella tularensis where AIM2-deficient mice
103863	AIM2	20457908	display an increased susceptibility to F. tularensis infection compared with wild-type mice. AIM2-containing inflammasomes are activated in response to cytosolic DNA, Â this response is augmented in keratinocytes from psoriatic lesions and
103863	AIM2	21562230	contributes to the auto-inflammatory disease.
103863	AIM2	25641891	Single nucleotide polymorphisms in IFI16 and AIM2 are associated with Behçet disease . IFI16, a PYHIN protein, is an intracellular DNA sensor that mediates the induction of interferon-beta (IFNB)
103852	IFI16	20890285	by directly associating with IFNB-inducing viral DNA motifs.

IFI16 acts as a nuclear pathogen sensor	and interacts			
with PYCARD and CASP1 to form	a functional			
inflammasome during KSHV infections.				

100050		01555000	inflammasome during KSHV infections.
103852	IFI16	21575908	
103852	IFI16	24131791	Cyclic-di-GMP-induced levels of IFI16 suppress the expression of TMEM173 (STING).
			IFI16 is essential for host defence by clustering into signalling foci with foreign DNA in a switch-like manner and is capable of using the size of naked double stranded DNA as a molecular ruler to distinguish self
103852	IFI16	24367117	from nonself. In herpes simplex virus 1 (HSV-1) infected cells, the stability and function of IFI16 and TMEM173 are dependent on cell derivation and the functional integrity of HSV-1 proteins ICP0 and US3 protein
103852	IFI16	24449861	kinase. IFI16 is a sensor for lentiviral reverse transcription
103852	IFI16	24154727	products and restricts HIV-1 replication in macrophages. IFI16 oligomerizes upon viral DNA sensing in human
103852	IFI16	24237704	cytomegalovirus (HCMV) infected-cells. The HCMV major tegument protein pUL83 blocks nuclear IFI16 oligomerization and inhibits IFI16-mediated antiviral cytokine expression The end result of the interplay between TMEM173 (STING), IFI16, and herpes simplex virus 1 (HSV-1) is
103852	IFI16	24449861	determined by the genotype of the infected cells and the functional integrity of HSV-1 proteins infected cell protein 0 (ICP0) and US3 protein kinase. IFI16 is a sensor for lentiviral reverse transcription products and restricts HIV-1 replication in human
103852	IFI16	24154727	macrophages. IFI16 transcriptionally regulates type-I interferons and
103852	IFI16	25002588	DDX58 (RIG-I) and controls the interferon response to both DNA and RNA viruses. IFI16 and MB21D1 interact and cooperate during
103852	IFI16	25831530	herpes simplex virus infection to initiate innate signalling. IFI16 restricts chromatinized human papillomaviruses
103852	IFI16	25972554	DNA through epigenetic modifications, thus reducing both viral replication and transcription. Single nucleotide polymorphisms in IFI16 and AIM2
103852	IFI16	25641891	are associated with Behçet disease. MAPK8 phosphorylates IRF3 and is essential for IRF3 dimerization induced by polyinosinic-cytidylic acid
73479	MAPK8	19153595	<ul><li>(polyI:C).</li><li>Mycobacterium tuberculosis phosphatase PtpA suppresses innate immunity by binding to ubiquitin; which, in turn, activates it to dephosphorylate</li></ul>
73479	MAPK8	25642820	phosphorylated MAPK8 and MAPK14.

84613	MAPK14	15569672	MAPK14 is dephosphorylated by PPP2R4, which then induces apoptosis in neutrophils and the resolution of inflammation. MAPK14 activation is blocked by Bacillus anthracis,
			resulting in the opening of a connexin ATP release channel and induction of macrophage death. Constitutive activation of MAPK14 interferes with inflammasome activation and IL1B production, which
84613	MAPK14	21683629	compromises antimicrobial immunity. (Demonstrated in murine model) The MAPK14 pathway is an important contributor to microglial production of proinflammatory cytokines induced by LPS or beta-amyloid. (Demonstrated in
84613	MAPK14	21733175	mouse) MAPK14 mediates cytoskeletal remodelling and early spreading of lipopolysaccharide (LPS)-stimulated
84613	MAPK14	22028692	macrophages. (Demonstrated in mice) MAPK1 (ERK) and MAPK14 (p38) control the
84613	MAPK14	22447027	dynamic balance regulating neutrophil migration. Mycobacterium tuberculosis phosphatase PtpA suppresses innate immunity by binding to ubiquitin; which, in turn, activates it to dephosphorylate
84613	MAPK14	25642820	phosphorylated MAPK8 and MAPK14. Pten is an essential regulator of natural killer cell localization in vivo during both homeostasis and
158878	Pten	25646418	malignancy. The and The and Th
135792	Tnfaip3	21220427	tolerance through the reprogramming of Tlr4 signaling.
			Tnfaip3 promotes intestinal epithelial barrier integrity and inhibits LPS-induced loss of the tight junction
135792	Tnfaip3	22031828	protein occludin. Tnfaip3 is regulated by both Nf-ΰB and p38-dependent
135792	Tnfaip3	24023826	Cebpb in response to LPS in macrophages. Mirlet7f and its target Tnfaip3 regulate immune responses to Mycobacterium tuberculosis and control bacterial burden by augmenting the production of Tnf
135792	Tnfaip3	25683052	and II1b. Sqstm1 captures Tnfaip3, an NFkB inhibitor, and sequesters it in the autophagosome. This allows macrophages to release chemokines that recruit
135792	Tnfaip3	25609235	neutrophils and boost antifungal immunity. Sqstm1 is required for Tlr4-mediated autophagy. Tlr4- driven induction of Sqstm1 plays an essential role in the formation and the autophagy degradation of aggresome-like induced structures, which might be critical for regulating host defense.
168358	Sqstm1	21220332	erritear for regulating nost defense.

Sqstm1 and Calcoco2 are ubiquitin-autophagy receptors that are required for the recognition of extracelluar bacterial DNA by the Tmem173 (STING)dependent cytosolic pathway, marking bacteria with ubiquitin, and delivery of bacilli to autophagosomes.

Sqstm1 Phosphorylation of Sqstm1 (p62) activates the Keap1-Nrf2 pathway during selective autophagy. 168358 Sqstm1 24011591 Sqstm1 captures Tnfaip3, an NFkB inhibitor, and sequesters it in the autophagosome. This allows macrophages to release chemokines that recruit 168358 Sqstm1 25609235 neutrophils and boost antifungal immunity. TREM1 is a superimmunoglobulin receptor present on neutrophils and monocytes, which plays an important role in the amplification of inflammation and its expression is inhibited by PGD(2) and PGJ(2) in 86629 TREM1 20797396 macrophages. TREM1 is an activating receptor expressed on neutrophils and monocytes that amplifies inflammation induced by TLR4-signalling, specifically in the 86629 TREM1 21393102 induction of TNFA production. TREM1 expression is upregulated following Pseudomonas aeruginosa infection in the cornea and the inhibition of TREM1 reduces the severity of corneal disease. TREM1 acts as an inflammatory amplifier in P. aeruginosa keratitis by modulating TLR signalling and Th1/Th2 responses. 86629 TREM1 21555403 TREM1 expression is induced by vitamin D3 in human bronchial epithelial cells. Activation of TREM1 leads to the induction of human beta defensin 2 and TNF-86629 TREM1 21690199 alpha mRNA in the airway epithelium. TREM1 mediates endotoxin tolerance in monocytes through its ability to induce anti- or pro-inflammatory 86629 TREM1 22459945 signals depending on its membrane-bound state. TREM1 expression and shedding are regulated by CpG-mediated TLR9 activation in macrophages. 86629 TREM1 23475790 PGLYRP1 binds to bacterially derived peptidoglycan and these complexes constitute potent ligands capable of binding to TREM1 and inducing myeloid cell 86629 TREM1 25595774 functions. PGLYRP1 is a secreted innate immunity protein that is expressed in polymorphonuclear leukocytes and is conserved from insects to mammals, it recognizes bacterial peptidoglycan, and functions in antibacterial 58413 PGLYRP1 20418257 immunity and inflammation. PGLYRP1 is a peptidoglycan recognition protein and play a role in innate immunity against L. monocytogenes infection by inducing TNFA. (Demonstrated in murine model)

22901810

168358

58413

PGLYRP1

21134971

58413	PGLYRP1	21439073	PGLYRP1 is a member of the Peptidoglycan Recognition Proteins (PGRP) family and recognizes peptidoglycan, a structural component of bacterial cell walls, as a part of innate immune response against infections. PGLYRP1 binds to Gram-positive bacterial wall and activates a protein-sensing two-component system to induce bacterial death. PLGYRP1-mediated activation results in membrane depolarization and cessation of peptidoglycan, protein, and RNA/DNA synthesis, as
58413	PGLYRP1	21602801	well as the production of hydroxyl radicals.
58413	PGLYRP1	25595774	PGLYRP1 binds to bacterially derived peptidoglycan and these complexes constitute potent ligands capable of binding to TREM1 and inducing myeloid cell functions. Defb1 is important for the control of early mucosal
136612	Il17f	25595775	Candida infection and plays a critical role in the induction of innate inflammatory mediators including, II1b, II6, Cxcl1, II17a, and II17f. Defb1 is important for the control of early mucosal Candida infection and plays a critical role in the induction of innate inflammatory mediators including, II1b, II6, Cxcl1, II17a, and II17f.
178943	Cxcl1	25595775	Defb1 is upregulated in plasmacytoid dendritic cells and monocyte during viral challenge Defb1-deficient mice infected with mouse-adapted HK18 (influenza) lost weight earlier and died sooner than WT mice, suggesting that Defb1 plays a role in early innate immune responses against influenza in vivo. However, lung virus titers were equal between the two mouse strains, indicating that the mechanism is not related to viral replication.
139170	Defb1	21551252	Defb1 is a component of platelets that displays classic antimicrobial activity and signals polymorphonuclear
139170	Defb1	22102811	leukocytes to extrude DNA lattices that capture and kill bacteria. (Demonstrated in human) Defb1 is important for the control of early mucosal Candida infection and plays a critical role in the induction of innate inflammatory mediators including,
139170	Defb1	25595775	II1b, II6, Cxcl1, II17a, and II17f.
175257	Dnase2a	25600358	Dnase2a is required for Tlr9 activation by bacterial genomic DNA. FPR1 and FPR2 are G-protein-coupled receptors that recognize bacterial signal peptides, constituting a novel
66320	FPR2	25605714	class of immune activators that contribute to mammalian immune defence against bacteria.

			FPR1 and FPR2 are G-protein-coupled receptors that recognize bacterial signal peptides, constituting a novel class of immune activators that contribute to
66270	FPR1	25605714	mammalian immune defence against bacteria. PPARG functions as an antimicrobial factor by
18954	PPARG	20421464	maintaining constitutive epithelial expression of a subset of beta-defensin in the colon. PPARG negatively regulates IFNB production in TLR3/4-stimulated macrophages by preventing IRF3 binding to the IFN-beta promoter.
18954	PPARG	21148557	MIR130A reduces hepatitis B virus (HBV) replication by down-regulating the expression of two major
18954	PPARG	25595716	metabolic regulators PPARGC1A and PPARG, both of which can potently stimulate HBV replication. PPARGC1A is activated in Staphylococcus aureus- mediated sepsis via the TLR2-signalling pathway.
11756	PPARGC1A	21966468	(Demonstrated in mice) MIR130A reduces hepatitis B virus (HBV) replication by down-regulating the expression of two major
11756	PPARGC1A	25595716	<ul> <li>metabolic regulators PPARGC1A and PPARG, both of which can potently stimulate HBV replication.</li> <li>MIR130A reduces hepatitis B virus (HBV) replication by down-regulating the expression of two major metabolic regulators PPARGC1A and PPARG, both of</li> </ul>
126561	MIR130A	25595716	which can potently stimulate HBV replication. Super-low dose lipopolysaccharide induces inhibitory phosphorylation of Pik3c3 leading to the disruption of endosome-lysosome fusion and low-grade
212409	Tollip	25586187	inflammation in innate macrophages; a mechanism that depends on the clearance and relocation of Tollip. Class III phosphatidylinositol 3-kinases (PI3K) are required for downstream Arf6 regulation of CpG
132410	Pik3c3	22170068	oligodeoxynucleotide uptake and thus have a role in Tlr9-mediated immune signalling. Super-low dose lipopolysaccharide induces inhibitory phosphorylation of Pik3c3 leading to the disruption of endosome-lysosome fusion and low-grade
132410	Pik3c3	25586187	inflammation in innate macrophages; a mechanism that depends on the clearance and relocation of Tollip. The pre-treated macrophages exhibit endotoxin tolerance, i.e. less cytokine production, towards LPS challenge. Thermediated cross-tolerization is mediated by suppression of LPS-induced signalling and
177358	Tnf	21602809	chromatin remodelling.
177358	Tnf	21611132	The sessential to mount an acute inflammatory response to dsDNA in the endothelium. Mirlet7f and its target The The sessent and control bacterial burden by augmenting the production of The sessent acute and the sessent acute and the sessent acute
177358	Tnf	25683052	and II1b.

			Cav1 is an important component of the innate host immune response to the majority of non-cytotoxic strains of P. aeruginosa by promoting bacterial clearance during acute pneumonia and chronic
129000	Cavl	19949109	colonization. The Akt1/Mir199a/Cav1 pathway is a regulator of innate immunity that is dysfunctional in cystic fibrosis
129000	Cav1	25665524	macrophages contributing to lung hyper-inflammation. The Akt1/Mir199a/Cav1 pathway is a regulator of
223741	Mir199a-1	25665524	innate immunity that is dysfunctional in cystic fibrosis macrophages contributing to lung hyper-inflammation. Akt1 is a Ser/Thr protein kinase that plays a pivotal role in functional activation in macrophages. Akt1 specifically functions in phagocytosis, intracellular bacterial infection, LPS tolerance, production of inflammatory cytokines/mediators, and migration during macrophage-mediate innate immunity.
174623	Akt1	21196185	daring maerophage mediate innate initiality.
			Akt1 activation is blocked by Bacillus anthracis, resulting in the opening of a connexin ATP release channel and induction of macrophage death. Constitutive activation of Akt1 interferes with inflammasome activation and Il1b production, which
174623	Akt1	21683629	compromises antimicrobial immunity. Akt1 functions downstream of Tlr2-stimulation to induce the expression of the monocyte chemoattractant
174623	Akt1	22218715	protein 1, Ccl2. Lysophosphatidic acid plays an anti-inflammatory role in macrophages by diminishing lipopolysaccharide- induced phosphorylation of Mapk14 and Akt1, as well
174623	Akt1	25783839	as Rela nuclear translocation. The Akt1/Mir199a/Cav1 pathway is a regulator of innate immunity that is dysfunctional in cystic fibrosis
174623	Akt1	25665524	macrophages contributing to lung hyper-inflammation. DR1, a novel host susceptibility gene for influenza A
100329	DR 1	25589657	<ul> <li>virus replication, suppresses host innate immunity and enhances viral RNA replication.</li> <li>II10 contributes to antiviral innate immunity during acute infection by restricting activation-induced death in natural killer (NK) cells. Blockade of II10 receptor during acute murine cytomegalovirus (CMV) infection</li> </ul>
190697	1110	21849677	markedly reduced the accumulation of cytotoxic NK cells in the spleen and lung. Il10 has opposing functions in anti-microbial responses in its capacity to mediate protective immunity against some organisms but increase susceptibility to other
190697	1110	22268692	infections. Il10-mediated suppression of natural killer/dendritic cell crosstalk leads to prolonged mouse
190697	II10	22876184	cytomegalovirus (MCMV) persistence due to poor priming of MCMV-specific T cells.

			Retinoic acid treatment enhances Tlr2-dependent II10 production from T cells and this, in turn, potentiates T
190697	II10	25826367	regulatory cell generation without the need for activation of antigen presenting cells. Intestinal macrophages that constitutively produce IL10, control excessive innate immune activation and
190697	II10	25959063	prevent tissue damage after an acute bacterial infection. CASP4 is a critical regulator of noncanonical inflammasome activation that initiates defence against
69557	CASP4	25964352	bacterial pathogens in primary macrophages by mediating cell death and IL1A release PTX3 is a multifunctional soluble molecule involved in inflammation and innate immunity. It is a unique factor H (FH) ligand in that it can bind both of the two hot-spots of FH and can participate in the localization
62945	PTX3	19050261	of functionally active FH. PTX3, an essential component of humoral innate immunity, and immunoglobulins share functional outputs, including complement activation, opsonization and glycosylation-dependent regulation of
62945	PTX3	20208538	inflammation. PTX3, an inflammation-associated long pentraxin, plays key roles in innate immunity, female fertility and vascular biology. PTX3 octamer contains two fibroblast growth factor 2 (FGF2) binding sites and this
62945	PTX3	20363749	quaternary organization is required for the anti- angiogenic function of PTX3. PTX3 is produced by innate immunity cells (e.g. PMN, macrophages, dendritic cells) and it interacts with several ligands to play an essential role in innate immunity, tuning inflammation and matrix deposition.
62945	PTX3	20683616	<ul> <li>PTX3 provides a paradigm for the mode of action of humoral innate immunity.</li> <li>PTX3, serum amyloid P component (SAP) and C-reactive protein (CRP) belong to the pentraxin family of pattern recognition molecules involved in tissue homeostasis and innate immunity. PTX3 heterocomplexes with mannose-binding lectin (MBL)</li> </ul>
62945	PTX3	21106539	to trigger cross-activation of the complement system. PTX3 production is up-regulated in response to serum amyloid A, and contributes to the inflammatory
62945	PTX3	21465531	pathogenesis of atherosclerosis. FCN3 and PTX3 are soluble oligomeric pattern- recognition molecules that interact with each other and
62945	PTX3	21490156	act synergistically to activate the lectin complement pathway. PTX3 forms a complex with components of neutrophil
62945	PTX3	22278372	extracellular traps in septic patients. Sputum PTX3 level are lower in cystic fibrosis patients
62945	PTX3	23475792	due to proteolytic cleavage by Aspergillus fumigatus.

			PTX3 acts as an extrinsic oncosuppressor gene by
62945	PTX3	25679762	regulating Complement-dependent, macrophage- sustained, tumor-promoting inflammation. PTX3 plays a non-redundant protective role in
62945	PTX3	25964372	orchestrating tissue repair and remodeling by interacting with fibrin and plasminogen. S100A12 is induced in response to H. pylori infection
102649	S100A12	25964473	<ul><li>and inhibits bacterial growth and viability in vitro by</li><li>binding nutrient zinc.</li><li>CXCL14 is important for constitutive antimicrobial</li></ul>
45673	CXCL14	25964486	defences against pneumonia Myo18a isoforms differentially regulate trafficking,
201810	Myo18a	25965346	expression, and activation of innate immune receptors on macrophages Cd81 inhibits Rac1/Stat1 activation and negatively
207763	Rac1	25972472	regulates the defence mechanisms to Listeria monocytogenes infection. Cd81 inhibits Rac1/Stat1 activation and negatively
212677	Cd81	25972472	regulates the defence mechanisms to Listeria monocytogenes infection.
			The G3BP1-CAPRIN1-PRKRA complex represents a new mode of PRKRA activation and links stress responses with innate immune activation through
38977	CAPRIN1	25784705	PRKRA without a requirement for foreign double- stranded RNA pattern recognition. EIF2AK2 (PKR) is recruited to stress granules by
54619	G3BP1	25520508	G3BP1 to promote innate immune responses at both transcriptional and translational levels. The G3BP1-CAPRIN1-PRKRA complex represents a
			new mode of PRKRA activation and links stress responses with innate immune activation through
54619	G3BP1	25784705	PRKRA without a requirement for foreign double- stranded RNA pattern recognition. Stimulation of TMEM173-dependent IRF3 activation
( = ) = 2			by ultraviolet radiation is due to apoptotic signalling- dependent disruption of ULK1, a pro-autophagic
65053	ULK1	25792739	protein that negatively regulates TMEM173. Nfil3 is an Ill2b transcriptional inhibitor in macrophages. Interactions of macrophages with the
152923	Nfil3	21383239	enteric microbiota induce Nfil3 to limit their inflammatory capacity. Nfil3 is a key regulator of common helper-like innate
152923	Nfil3	25801035	lymphoid cell progenitors as they emerge during early lymphopoiesis. Lipopolysaccharide-mediated myeloid Anpep (CD13)
			expression governs internalization of Tlr4 and negatively regulates Tlr4 signalling, thereby balancing
194391	Anpep	25801433	the innate response by maintaining the inflammatory equilibrium critical to innate immune regulation. WFDC12 plays a role in the regulation of lung
77255	WFDC12	25770093	inflammation.

184766	Ctnnd1	25773174	Ctnnd1 expressed in alveolar type II epithelial cells plays a critical role in regulating the innate immunity of the entire lung.
150505		25701027	Escherichia coli toxin CNF1 promotes the maturation/secretion of II1b while the $\hat{I} \pm$ -hemolysin toxin inhibits II1b secretion without affecting the
150795	Ly6g	25781937	recruitment of Ly6g+ cells. Rela is a subunit of NFKB and is not essential for virus-stimulated Ifnb expression, instead, Rela sustains autocrine Ifnb signaling prior to infection. The absence of Rela causes significant delays in Ifnb induction and consequently defective secondary antiviral gene expression. Rela maintains autocrine Ifnb signaling in uninfected cells, facilitates inflammatory and adaptive immune responses following infection, and promotes infected cell survival during this process.
132298	Rela	21209118	
			Rela is critical for pulmonary host defense during Streptococcus pneumoniae pneumonia in alveolar macrophages. During pneumococcal pneumonia, only the earliest induction of cytokines depends on transcription regulated by Rela in myeloid cells, and this transcriptional activity contributes to effective immunity.
132298	Rela	21216972	Rela is required for Il17a production in T cell in response to bacterial infection. Rela deficient T cells resulted in a diminished innate immune response to E.
132298	Rela	21419662	coli infection. Tnfaip3 is regulated by both Nf-ΰB and p38-dependent
132298	Rela	24023826	Cebpb in response to LPS in macrophages. The noncanonical NF ΰB pathway regulates histone modifications at the Ifnb1 promoter resulting in attenuated recruitment of Rela and histone demethylase, Kdm4a, to the Ifnb1 promoter. This provides a mechanism for regulating the induction of
132298	Rela	24656046	type I interferons . Lysophosphatidic acid plays an anti-inflammatory role in macrophages by diminishing lipopolysaccharide- induced phosphorylation of Mapk14 and Akt1, as well
132298	Rela	25783839	as Rela nuclear translocation. IL1RL1 (ST2) pre-treatment suppresses cytokine production and inhibits LPS signalling in dendritic
148423	Il1rl1	22922442	cells. (Demonstrated in human) Cigarette smoke decreases Il1rl1 expression on group 2 innate lymphoid cells while elevating Il1rl1 expression on macrophages and natural killer cells, thus altering
148423	Il1rl1	25786179	Il33 responsiveness within the lung to infection.

			HAVCR2 is constitutively expressed on human resting monocytes/macrophages and functions as a cap to block IL12, which is a key pro-inflammatory cytokine linking innate and adaptive immune responses. HAVCR2 plays a crucial role in the negative regulation of innate immune responses through crosstalk with DDCD1 and SOCS1 to limit STAT1 phasehomilation
55271	HAVCR2	21637332	<ul> <li>PDCD1 and SOCS1 to limit STAT1 phosphorylation in HCV infection.</li> <li>TLR activation promotes HAVCR2 and LGALS9 association within the same macrophage to differentially regulate IL12/IL23 expressions via</li> </ul>
55271	HAVCR2	23967307	differentially regulate IL12/IL23 expressions via STAT3 phosphorylation. Hepatitis C virus (HCV)-induced, MIR155-regulated HAVCR2 expression regulates natural killer cell function, suggesting a novel mechanism for balancing immune clearance and immune injury during chronic
55271	HAVCR2	25772938	viral infection. HIF1A transcription is induced by IFNA2 in human endothelial cells though a JAK-ISGF3 pathway under normoxic conditions, and that this response contributes
8510	HIF1A	18606657	to the anti-proliferative activity of this cytokine. HIF1A expression is regulated by intracellular calcium levels, resulting in modulation of PPP3CA
8510	HIF1A	17965024	<ul> <li>(calcineurin) activity and RACK1 dimerization.</li> <li>Hypoxia-inducible factors (HIFs), including HIF1A, regulate glycolytic energy generation, optimize innate immunity, control pro-inflammatory gene expression, mediate bacterial killing and influence cell migration.</li> <li>HIFs contribute to inflammatory functions in various</li> </ul>
8510	HIF1A	20517715	components of innate immunity, such as neutrophils, dendritic cells, mast cells, and epithelial cells. HIF1A, the most ubiquitously expressed hypoxia- inducible factor (HIF), in epithelial cells alters the
8510	HIF1A	20511350	lung's innate immune response and biases the tissue toward a Th2-mediated inflammation. HIF1A, under normoxic conditions, accumulates in dendritic cells via the TLR/MYD88/NFkB signalling pathway to induce a distinct subset of proinflammatory
8510	HIF1A	21685248	<ul><li>genes in comparison to hypoxia-induced HIF1A.</li><li>(Demonstrated in murine model)</li><li>HIF1A mediates the functional plasticity of monocytes</li><li>during sepsis, wherein they transit from a pro- inflammatory to an immunosuppressive phenotype,</li></ul>
8510	HIF1A	25746953	while enhancing protective functions like phagocytosis, anti-microbial activity, and tissue remodelling . CHUK is part of the inhibitor kappaB kinase (IKK) complex that phosphorylates IRS1 at Ser(312) and this
243385	CHUK	12351658	contributes to the insulin resistance mediated by activation of inflammatory pathways. CHUK (IKK-alpha) is part of the IKK signalosome that phosphorylates NFKBIA and NFKBIB, leading to
243385	СНИК	9891086	activation of NF-kappaB.

			CHUK (IKKalpha) phosphorylates CREBBP (CBP), regulating the CBP-mediated crosstalk between NF-kappaB and p53, a critical factor in the promotion of
243385	CHUK	17434128	cell proliferation and tumor growth. CHUK is a unique molecule involved in TLR7/9-
243385	CHUK	20200270	<ul> <li>MyD88-dependent type I IFN production through dendritic cell subset-specific mechanisms.</li> <li>CHUK has a key role in the negative feedback of NF-kB canonical signalling by orchestrating the assembly of the A20 ubiquitin-editing complex to limit inflammatory gene activation in response to proinflammatory stimuli such as TNF and IL1.</li> </ul>
243385	CHUK	21765415	(Demonstrated in mouse) CHUK is required in dendritic cells to prime adaptive
243385	СНИК	23422957	immunity to Listeria monocytogenes. GNB2L1 (RACK1) negatively regulates NF $\hat{I} \circ B$ activation by interacting with CHUK and IKBKB. The interaction interferes with the recruitment of the IKK
243385	СНИК	24323043	complex to TRAF2. DDX3X initiates a multifaceted cellular program involving dynamic associations with hepatitis C virus (HCV) RNA and proteins, CHUK, stress granules, and lipid droplet surfaces for its crucial role in the HCV life
243385	СНИК	25740981	cycle. Ptpn11 phosphatase function positively regulates Clec7a- and Itgam-stimulated reactive oxygen species production in macrophages by dephosphorylating and thus mitigating the inhibitory function of Sirpa and by
209781	Mapk3	25538234	promoting Mapk1/Mapk3 activation. Prostaglandin E2 transactivates Csf1r and synergizes with its signalling at Mapk1/Mapk3 level in promoting
209781	Mapk3	25757564	macrophage migration. Mapk1 (ERK) and Mapk14 (p38) control the dynamic balance regulating neutrophil migration.
137672	Mapk 1	22447027	(Demonstrated in human) Ptpn11 phosphatase function positively regulates Clec7a- and Itgam-stimulated reactive oxygen species production in macrophages by dephosphorylating and thus mitigating the inhibitory function of Sirpa and by
137672	Mapk1	25538234	promoting Mapk1/Mapk3 activation. Prostaglandin E2 transactivates Csf1r and synergizes with its signalling at Mapk1/Mapk3 level in promoting
137672	Mapk1	25757564	macrophage migration. Prostaglandin E2 transactivates Csf1r and synergizes with its signalling at Mapk1/Mapk3 level in promoting
150998	Csflr	25757564	macrophage migration. Traf2 deficiency results in the accumulation of Tnf- dependent, Il10-secreting neutrophils. Combined treatment of neutralizing antibodies against both Tnf and Il10 substantially ameliorated the colitis phenotype
146254	Traf2	22546736	in the Traf2 null mice.

			The binding of MAVS to Traf2, Traf5, and Traf6 is dependent on virus infection and MAVS polymerization . The TRAF proteins promote
146254	Traf2	23951545	ubiquitination that recruits IKBKG binding to the MAVS signalling complex. Traf2 mediates proteasome-dependent degradation of Irf5 and Rel as well as regulating macrophage
146254	Traf2	25565375	polarizationintumourmicroenvironmentandcontrolling tumour growth.Traf2-deficient macrophages produce reduced levels ofinflammatorycytokinesinresponseto
146254	Traf2	25752829	lipopolysaccharide or flagellin stimulation and exhibit increased susceptibility to S. Typhimurium infection. Cfp plays a role in intestinal homeostasis in response to an infectious challenge to activate Hc (C5a), which in
135734	Cfp	25725105	turn provides protection through Il6 expression by the epithelium. Cfp plays a role in intestinal homeostasis in response
163835	Нс	25725105	to an infectious challenge to activate Hc (C5a), which in turn provides protection through Il6 expression by the epithelium. IL18 and IL1B are important pro-inflammatory
71390	IL18	20195505	cytokines that activates monocytes, macropages, and neutrophils, as well as induce the Th1 and Th17 adaptive cellular responses. IL18 secreted by inflammatory monocytes is critical for
71390	IL18	22940097	<ul> <li>the differentiation of memory CD8(+) T and NK</li> <li>lymphocytes into antimicrobial effector cells.</li> <li>(Demonstrated in mice)</li> <li>Primary Î<sup>3</sup>Î' T cells provide an early source of IFNG</li> <li>during dengue virus (DV) infection and target DV-</li> </ul>
71390	IL18	25732728	infected cells. Monocytes also participate as accessory cells that sense DV infection and amplify the cellular immune response in an IL18-dependent manner. Tlr3 deletion dramatically enhanced the development of elastic lamina damage after collar-induced injury,
152685	Tlr3	21220319	indicating that Tlr3 signalling plays a protective role in arterial vessel wall. Tlr3 activation by Poly(I:C) in the endothelial cells
152685	Tlr3	21367858	induces Poly(I:C) dose- and time-dependent cell apoptosis. Specifically, Tlr3 stimulation triggered the signalling of both extrinsic and intrinsic apoptotic pathways. Tlr3 requires proteolytic processing in endolysosome
152685	Tlr3	21402738	by asparagine endopeptidase and cathepsin in the endolysosome to initiate signalling in response to DNA. Tlr3 expression is inducible by LPS via Tlr4-Myd88-
152685	Tlr3	21498625	Irak-Traf6-NfkB dependent signalling pathway. (Demonstrated in human)

			Tlr3 is necessary to establish an antiviral state in hepatocytes infected with hepatitis C Virus. HCV envelope proteins counteract the antiviral host defence by inhibiting the expression of Tlr3. (Demonstrated in
152685	Tlr3	21695051	human) TLR3 signalling is enhanced by the presence of viral
152685	Tlr3	22016778	double-strand RNA-binding proteins. (Demonstrated in human) Tlr3-Ticam1-mediated signalling pathway plays an
152685	Tlr3	22072781	essential role in the anti-viral response against poliovirus infection. Tlr3 is constitutively expressed in spermatogonia and
152685	Tlr3	22262694	spermatocytes, and has the ability to activate anti-viral responses. Upon engagement with its ligand, dsRNA, Tlr3
152685	Tlr3	22421964	possesses the ability to recruit Casp8 and Ripk1 to induce apoptosis. (Demonstrated in human) Activation of Tlr3 with poly(I:C) mediates antiviral
152685	Tlr3	22754655	immunity that diminishes coronavirus production in macrophages. Upregulation of Tlr3 in intestinal epithelia during
152685	Tlr3	22570612	infancy may contribute to age-dependent susceptibility to rotavirus infection. TLR3 activation differentially regulates phagocytosis
152685	Tlr3	22986631	of bacteria and apoptotic neutrophils by peritoneal macrophages.
152685	Tlr3	23142781	Tlr3 contributes to the control of activated endogenous retroviruses (ERVs) and ERV-induced tumours. Wdfy1 is a crucial adaptor protein in the Tlr3/4 signalling pathway. Wdfy1 interacts with Tlr3 and Tlr4
152685	Tlr3	25736436	and mediates the recruitment of Ticam1 to these receptors. Wdfy1 is a crucial adaptor protein in the Tlr3/4 signalling pathway. Wdfy1 interacts with Tlr3 and Tlr4
172895	Wdfy1	25736436	and mediates the recruitment of Ticam1 to these receptors.
157290	Earl1	25713137	Ear11 has prominent chemoattractant activity for F4/80(+)CD11c(-) tissue macrophages. Tnfrsf13c (BAFFR) expression is critical for innate immune activation and antiviral immunity. Tnfrsf13c deficiency results in reduced enforced viral replication,
166389	Tnfrsf13c	25673724	limited type I interferon production, and reduced adaptive immunity. Ptx3 production is up-regulated in response to serum amyloid A, and contributes to the inflammatory
149887	Ptx3	21465531	pathogenesis of atherosclerosis. (Demonstrated in human) Ptx3 forms a complex with components of neutrophil
149887	Ptx3	22278372	extracellular traps in septic patients. (Demonstrated in human)
			Ptx3 acts as an extrinsic oncosuppressor gene by regulating Complement-dependent, macrophage-
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149887	Ptx3	25679762	sustained, tumor-promoting inflammation. Stat3 is required for maximal Osm-induced lung Cxcl5
142284	Osm	25692402	expression and subsequent neutrophil recruitment during bacterial pneumonia. Stat3 is required for maximal Osm-induced lung Cxcl5
178768	Cxcl5	25692402	expression and subsequent neutrophil recruitment during bacterial pneumonia. Stat3 is a negative regulator of type I IFN-mediated
211637	Stat3	21810606	anti-viral responses. STAT3 mediates protein transport and secretion in
211637	Stat3	23460517	hepatocytes during the acute phase of pneumonia. Stat3 mediates protection against intestinal infection by
211637	Stat3	24412612	inducing innate lymphoid cell derived-II22. Stat3 is required for maximal Osm-induced lung Cxcl5
211637	Stat3	25692402	expression and subsequent neutrophil recruitment during bacterial pneumonia. Unrepaired DNA lesions induce type I interferons via the Tmem173 pathway, resulting in enhanced anti-viral
165511	Atm	25692705	and anti-bacterial responses in Atm (-/-) mice.
71032	CXCR4	21094463	CXCR4 is a chemokine receptor that is essential for homing of stem cells and more mature neutrophils to the bone marrow. MIR146A upregulation by CXCR4 antagonist
71032	CXCR4	25705792	<ul> <li>AMD3100 treatment or ZBTB16 silencing, decreases</li> <li>CXCR4 protein expression and prevents HIV-1</li> <li>infection of leukemic monocytic cell line and CD4(+)</li> <li>T lymphocytes.</li> <li>MIR146A upregulation by CXCR4 antagonist</li> <li>AMD3100 treatment or ZBTB16 silencing, decreases</li> <li>CXCR4 protein expression and prevents HIV-1</li> </ul>
72004	ZBTB16	25705792	infection of leukemic monocytic cell line and CD4(+) T lymphocytes. IL37 acts as an extracellular cytokine to inhibit innate
53480	IL1RAPL1	25654981	inflammation by binding to IL18R1 and then using IL1RAPL1 for its anti-inflammatory properties. IL37 acts as an extracellular cytokine to inhibit innate
64199	IL18R1	25654981	inflammation by binding to IL18R1 and then using IL1RAPL1 for its anti-inflammatory properties. IL1F7 (IL-37) is a natural suppressor of innate inflammatory and immune responses. IL1F7 expression in macrophages or epithelial cells suppresses production of pro-inflammatory cytokines
66559	IL1F7	20935647	and the abundance of these cytokines increases with silencing of endogenous IL1F7 in human blood cells.

			IL1F7 functions as a key modulator of intestinal inflammation where transgenic mice expressing IL1F7 subjected to dextran sulfate sodium-induced colitis showed reduced inflammation which was associated
66559	IL1F7	21873195	with decreased leukocyte recruitment into the colonic lamina propria. (Demonstrated in murine model) The precursor and mature forms of IL37 are secreted from activated cells upon inflammasome activation and
66559	IL1F7	24481253	<ul><li>CASP1 processing of IL37 is important for its anti- inflammatory activity.</li><li>IL37 acts as an extracellular cytokine to inhibit innate inflammation by binding to IL18R1 and then using</li></ul>
66559	IL1F7	25654981	IL1RAPL1 for its anti-inflammatory properties. Sf3a1 affects the mRNA splicing of genes in the TLR signalling pathway to modulate the innate immune
141954	Sf3a1	25658809	response in macrophages. CLEC6A (Dectin-2) recognition of alpha-mannans and induction of Th17 cell differentiation is essential for
17145	CLEC6A	20493731	host defence against Candida albicans. CLEC6A, a C-type lectin receptor, is a pattern recognition receptor critical for immune responses to fungi. CLEC6A is coupled to SYK kinase and signals via CARD9 to activate NFKB, which in turns induces both innate and adaptive immunity.
17145	CLEC6A	21267996	CLEC6A is critical for the development of house dust mite (Dermatophagoides farinae) elicited eosinophilic and neutrophilic pulmonary inflammation. CLEC6A was also found to be crucial for the Th2 cytokine
17145	CLEC6A	21357742	induction in the lungs and re-stimulated lymph nodes. CLEC6A is expressed mainly in DCs and macrophages. CLEC6A recognizes alpha-mannans with its carbohydrate recognition domain and transduces signals through association with the ITAM- containing Fc receptor gamma chain, which recruits
17145	CLEC6A	21677049	SYK and initiates the CARD9/NFkB signalling cascade. CLEC4D and CLEC6A form a heterodimer complex which confers innate cells high ability to sense C.
17145	CLEC6A	23911656	albicans infection by facilitating the activation of NFΰB dependent inflammatory responses. Plasmacytoid dendritic cells (pDCs) directly recognize Aspergillus fumigatus hyphae via CLEC6A; this
17145	CLEC6A	25659141	interaction results in antifungal activity and the formation of pDC extracellular traps. Cd200-Cd200r1 pathway is an important regulator of antiviral immunity during cytomegalovirus infection
162436	Cd200r1	25654642	that is exploited by murine cytomegalovirus to establish chronicity within mucosal tissue.

			Cd200-Cd200r1 pathway is an important regulator of antiviral immunity during cytomegalovirus infection
162936	Cd200	25654642	that is exploited by murine cytomegalovirus to establish chronicity within mucosal tissue. Stimulation of P2RX7 receptors activates ANO6 to enhance bacterial phagocytosis and killing by
28318	ANO6	25651887	macrophages. P2X7 activity is modulated by human cathelicidin CAMP (LL-37) receptor in a structure-dependent
61253	P2RX7	18765670	<ul><li>manner and is involved in the proliferative cell response to CAMP.</li><li>P2RX7 activation in lipopolysaccharide (LPS)-primed myeloid cells results in secretion of pro-inflammatory cytokines IL1B and IL18. In addition, P2RX7</li></ul>
61253	P2RX7	21988719	functions in the recognition and phagocytosis of non- opsonized bacteria and apoptotic cells. P2RX7 engagement with ATP analog initiates
61253	P2RX7	23479230	cutaneous inflammation, dendritic cell differentiation, and induction of Th17 immunity. Stimulation of P2RX7 receptors activates ANO6 to enhance bacterial phagocytosis and killing by
61253	P2RX7	25651887	macrophages. The transcription factor Zfp423 is necessary for adipocyte activation and impaired adipogenesis is
178353	Zfp423	25554785	observed in Zfp423(nur12) mice. Traf2 mediates proteasome-dependent degradation of Irf5 and Rel as well as regulating macrophage
155648	Rel	25565375	polarization in tumour microenvironment and controlling tumour growth. Phosphorylation of Irf5 at ser451 and ser462 is the primary trigger of Irf5 function in nuclear
132026	Irf5	22412986	accumulation, transcription and apoptosis. (Demonstrated in human) Irf5 modulates West Nile virus pathogenesis and host immune responses by shaping the early pro-
132026	Irf5	25031348	inflammatory cytokine response in the draining lymph node. Traf2 mediates proteasome-dependent degradation of
132026	Irf5	25565375	Irf5 and Rel as well as regulating macrophage polarization in tumour microenvironment and controlling tumour growth. SLC22A3 deficiency significantly decreases lipopolysaccharide- induced monocytic inflammatory
98707	SLC22A3	25561729	response by interrupting NF-kB and MAPKs (mitogen- activated protein kinases) signalling cascades in a histamine dependent manner. TPPII controls the balance between intracellular amino acid availability, lysosome number, and glycolysis,
48458	TPP2	25525876	which is vital for adaptive and innate immunity and neurodevelopmental health.

			EIF2AK2 is a protein kinase that is regulated by RNA and is an important mediator of the antiviral and anti
44867	EIF2AK2	12954221	proliferative actions of interferon (IFN). EIF2AK2 binds both SP1 and SP3 but only SP3 functions as part of the interferon-inducible complex with ISGF-3 proteins, consisting of STAT1, STAT2,
44867	EIF2AK2	12954221	and IRF9. EIF2AK2 enhances the induction of interferon-beta and
			apoptosis mediated by cytoplasmic RNA sensors where in addition to MAVS and IRF3 but not TRIF, it is required for maximal type I IFN-beta induction and the induction of apoptosis by both transfected T7 phage polymerase-synthesized RNAs (PRNAs) and
44867	EIF2AK2	19028691	polyinosinic-polycytidylic acid. EIF2AK2 is used by hepatitis C virus to restrain its ability to induce IFN through the RIG-I/MAVS
44867	EIF2AK2	20485506	pathway. Antiviral stress granules containing DDX58 (RIG-I) and EIF2AK2 (PKR) have a critical role in viral detection and innate immunity. (Demonstrated in
44867	EIF2AK2	22912779	mouse)
44867	EIF2AK2	23115300	Herpes simplex virus evades antiviral host defence by interacting with EIF2AK2 (PKR) to inhibit autophagy.
44867	EIF2AK2	23236554	Bacterial dsRNA or poly(I:C) induces tyrosine phosphorylation of PKR activation by JAK kinases. EIF2AK2 binds to dsRNA to inhibit IFN induction
44867	EIF2AK2	23372823	triggered by dengue virus. EIF2AK2 (PKR) is recruited to stress granules by
44867	EIF2AK2	25520508	G3BP1 to promote innate immune responses at both transcriptional and translational levels. RPS6KA5 negatively regulates TLR-pathway driven inflammation by preventing the binding of
16084	RPS6KA5	18690222	phosphorylated transcription factors CREB and ATF1 to IL10 and DUSP1 promoters.
16084	RPS6KA5	19922413	RPS6KA5 regulates the transcription of IL1RA in response to TLR activation in macrophages. Paramyxoviruses trigger the DNA-damage response, a pathway required for RPS6KA5 activation of phospho Ser 276 RELA formation to trigger the IRF7-DDX58
16084	RPS6KA5	25520509	amplification loop necessary for mucosal interferon production. IFITM3 is an antiviral restriction factor that mediates
16268	IFITM3	20064371	cellular resistance to influenza A H1N1 virus, West Nile virus, and Dengue virus. IFITM3 is post-translationally regulated by S- palmitoylation on membrane-proximal cysteines, which controls its clustering in membrane
16268	IFITM3	20601941	compartments and its antiviral activity against influenza virus.

			NRAV, a long noncoding RNA, modulates antiviral responses by negatively regulating the initial transcription of multiple critical interferon-stimulated
16268	IFITM3	25525793	genes, including IFITM3 and MX1, by affecting their histone modification. Upon cytoplasmic DNA stimulation, the endoplasmic
50271	INSIG1	25526307	reticulum protein AMFR is recruited to and interacts with TMEM173 in an INSIG1-dependent manner. Upon cytoplasmic DNA stimulation, the endoplasmic reticulum protein AMFR is recruited to and interacts
31785	AMFR	25526307	with TMEM173 in an INSIG1-dependent manner. Genetic deletion of Eif4ebp1 or Eif4ebp2 potentiates innate antiviral immunity by enhancing translation of
155976	Eif4ebp2	25531441	Irf7. Genetic deletion of Eif4ebp1 or Eif4ebp2 potentiates innate antiviral immunity by enhancing translation of
145947	Eif4ebp1	25531441	Irf7.
			Anxa1-/- mice are more susceptible to Mycobacterium tuberculosis infection as evidenced by a transient increase in pulmonary bacterial burden, exacerbated and disorganized granulomatous inflammation and impaired ability of Anxa1-/- dendritic cells to activate
151723	Anxal	25533809	naà ve T cells. Ptpn11 phosphatase function positively regulates Clec7a- and Itgam-stimulated reactive oxygen species production in macrophages by dephosphorylating and thus mitigating the inhibitory function of Sirpa and by
206000	Sirpa	25538234	promoting Mapk1/Mapk3 activation. Itgam (Cd11b integrin) is activated via Toll-like receptors (TLRs) and engages in crosstalk with the Myd88 and Ticam1 (TRIF) pathways inhibiting TLR
210531	Itgam	20639876	signalling in innate immune responses. Itgam :: Itgb2 is the principal leukocyte receptor involved in the recognition of the fungus Candida albicans. Recognition of Pra1p protein of C. albicans by Itgam :: Itgb2 plays a pivotal role in determining fungal virulence, and host response/protection against C. albicans infection.
210531	Itgam	21245270	Itgam (Cd11b) fine tunes the balance between adaptive and innate immune responses initiated by LPS by
210531	Itgam	24423728	modulating the trafficking and signalling functions of Tlr4 in a cell-type-specific manner. Ptpn11 phosphatase function positively regulates Clec7a- and Itgam-stimulated reactive oxygen species production in macrophages by dephosphorylating and thus mitigating the inhibitory function of Sime and by
210531	Itgam	25538234	thus mitigating the inhibitory function of Sirpa and by promoting Mapk1/Mapk3 activation.

0.5 ( 770		212(700)	Clec7a, a C-type lectin receptor, is a pattern recognition receptor critical for immune responses to fungi. Clec7a is coupled to Syk kinase and signals via Card9 to activate NFKB, which in turns induces both innate and adaptive immunity.
256772	Clec7a	21267996	Clec7a is expressed mainly in DCs and macrophages. Clec7a recognizes beta-glucans with its carbohydrate recognition domain and transduces signals through its ITAM-like motif in the cytoplasmic region, which recruits Syk and initiates the Card9/NFkB signalling
256772	Clec7a	21677049	cascade. Clec7a acts as an extracellular sensor for fungi and mycobacteria that induce both Il1b production and maturation for protective immunity. (Demonstrated in
256772	Clec7a	22267217	human) The functional activity of Clec7a in mucosal immunity to Candida albicans is dependent on the genetic background of the host; this was specifically observed
256772	Clec7a	22543832	in two strains of mice, C57BL/6 and BALB/c. Clec7a is necessary for macrophage activation and
256772	Clec7a	23386437	resistance to pathogenic fungus Coccidioidesimmitis. Ptpn11 phosphatase function positively regulates Clec7a- and Itgam-stimulated reactive oxygen species production in macrophages by dephosphorylating and
256772	Clec7a	25538234	thus mitigating the inhibitory function of Sirpa and by promoting Mapk1/Mapk3 activation. Ptpn11 phosphatase function positively regulates Clec7a- and Itgam-stimulated reactive oxygen species production in macrophages by dephosphorylating and
196834	Ptpn11	25538234	thus mitigating the inhibitory function of Sirpa and by promoting Mapk1/Mapk3 activation. IRAK4-mediated innate immune inflammatory responses play critical roles in divergent clinical outcomes in murine malaria models when Irak4 (-/-)
177236	Irak4	20595480	mice were used to study two experimental models of malaria. Irak4 activation is impaired during endotoxin tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of anti-inflammatory or anti- microbial mediators.
177236	Irak4	24717937	Irak4 deficient mice show impaired innate immunity, leading to defective Type 1 T-cell responses, B-cell
177236	Irak4	23027530	expansion, and are more susceptible to Toxoplasma gondii infection. Mir302b expression is up-regulated upon bacterial infection and is a crucial regulator of NFΰB signalling
177236	Irak4	24717937	by directly targeting IRAK4

			Irak4 is a molecular target of chlorogenic acid in the treatment of innate immunity-related shock and organ
177236	Irak4	25548221	dysfunction following insult of various Toll-like receptor pathogens from bacteria and viruses. Tmem126a upregulates genes involved in antigen
212533	Cd40	25549946	presentation; such as Icam1, MHC II, Cd86 and Cd40, via the Tlr4 signal transduction pathway. Tmem126a upregulates genes involved in antigen
158071	Cd86	25549946	presentation; such as Icam1, MHC II, Cd86 and Cd40, via the Tlr4 signal transduction pathway. Tmem126a upregulates genes involved in antigen
138668	Icam 1	25549946	presentation; such as Icam1, MHC II, Cd86 and Cd40, via the Tlr4 signal transduction pathway. Tmem126a upregulates genes involved in antigen
198859	Tmem126a	25549946	presentation; such as Icam1, MHC II, Cd86 and Cd40, via the Tlr4 signal transduction pathway. TP53 binds to canonical and non-canonical promoter regions of the human TLR gene family and up- regulates the expression of TLRs. The activation of
26364	TP53	21483755	TP53 can directly influence the TLR-mediated induction of cytokines. TP53 serves as a host antiviral factor and enhances
26364	TP53	22105999	both the innate and adaptive immune responses to influenza A virus. (Demonstrated in mice) Coronavirus engages papain-like proteases to escape
26364	TP53	25505178	from the innate antiviral response of the host by inhibiting TP53-IRF7-IFNB1 signalling.
178961	Atg7	25512546	Atg7 plays an essential role for autophagy during invariant natural killer T cell development. Arl5b negatively regulates the antiviral innate immune
138680	Arl5b	25451939	response by binding to Ifih1 and prevents the subsequent interaction of Ifih1 to poly(I:C).
282076	MIR548G	25499200	Overexpression of MIR548G suppresses multiplication and translation of dengue virus (DENV) 1, 2, 3 and 4. S100A8 is an intracellular calcium-binding protein that
102654	S100A8	12626582	promotes neutrophil/monocyte recruitment at inflamed tissues by enhancing attachment to endothelial cells. S100A8 is a myeloid-related protein that rapidly
102654	S100A8	18714033	<ul> <li>modulates macrophage nitric oxide production during innate immune response.</li> <li>S100A8 (calgranulin A) and S100A9 (calgranulin B) form an antimicrobial heterodimeric complex known as calprotectin. Bacterial flagellin induces the upregulation of S100A8/S100A9 heterodimer via a</li> </ul>
102654	S100A8	20555353	TLR5-dependent mechanism in epidermal keratinocytes. S100A8 forms a complex with S100A9 and the complex is the site of interplay between extracellular Ca(2+) entry and intra-phagosomal reactive oxygen species production. S100A8 :: S100A9 acts as Ca(2+) sensor in phagosomal ROS production.
102654	S100A8	21239714	

			S100A8 is strongly upregulated in neutrophils upon bacterial infection, and sequesters zinc as a mechanism of nutritional immunity. Salmonella typhimurium
102654	S100A8	22423963	overcomes this defence mechanism by expressing a high affinity zinc transporter. (Demonstrated in mice) S100A8::S100A9 heterodimer sequesters Mn(2+) and
102654	S100A8	23431180	Zn(2+) to starve bacteria of these essential nutrients. The TLR4/S100A8 axis is important in the activation
102654	S100A8	25505274	of monocytes. IL17A is a cytokine produced by T helper 17 (Th17)
90504	IL17A	19144317	cells that plays important roles in the development of inflammatory diseases. IL17A is an innate-adaptive immunomodulatory cytokine that is produced by gammadelta cells and is a key mediator for the innate immune response to urinary tract infections (UTIs) caused by uropathogenic
90504	IL17A	20083670	Escherichia coli. IL17A is a Th17-related cytokine, traditionally thought of as an adaptive responder, has been shown to have various innate sources and functions as a rapidly produced pro-inflammatory mediator. Innate IL17A- producing cells also employ many of the cytokine and
90504	IL17A	21074482	transcriptional regulators utilized by Th17 cells. IL17A signalling enhances the mRNA stability of
90504	IL17A	21822258	chemokine CXCL1 through TRAF3IP2, TRAF2- TRAF5 and the RNA-binding protein SRSF1. IL17A is significantly upregulated in both S. pyogenes inoculated and mock inoculated mice, indicating that the cytokine production can be triggered by inoculation
90504	IL17A	22384827	trauma alone. (Demonstrated in mice) Local production of IL17A in the airways drives early neutrophil infiltration into respiratory syncytial virus
90504	IL17A	24194936	infected infant lungs. IFNG interferes with the IL-1/NFKBIZ axis in $\hat{1}^2$ -glucan-activated dendritic cells and promotes T cell- mediated immune responses with increased release of
90504	IL17A	25474109	IFNG and IL22, and diminished production of IL17A. IL22 protects against and IL22RA2 aggravates liver
96992	IL22RA2	25476703	fibrosis and cirrhosis in chronic liver infections. Cxcl2 is a chemokine secreted by kidney dendritic cells to recruit neutrophils to sites of uropathogenic bacterial
178955	Cxcl2	21757770	infection. Cebpa suppresses granule formation in mast cells and
178955	Cxcl2	25447519	increases Cxcl2 production from mast cells upon bacterial stimulation. Cebpa is a member of the CCAAT enhancer binding protein family and is a transcriptional factor regulating genes in innate immunity and inflammation. The activities of CEBP are regulated via methylation of
173713	Cebpa	21326902	arginine and lysine side chains.

173713	Cebpa	25447519	Cebpa suppresses granule formation in mast cells and increases Cxcl2 production from mast cells upon bacterial stimulation. TLR6 co-expressed withTLR2 at the cell surface is crucial for recognition of diacylated lipopeptide and peptidoglycan derived from mycoplasma and to
13675	TLR6	15661917	activate the NF-kappaB signalling cascades in human cells. TLR6 and TLR1 are involved in the discrimination of a
13675	TLR6	12697090	subtle difference between triacyl and diacyl lipopeptides through interaction with TLR2. TLR6/2 heterodimer signalling is used by CD36, a
13675	TLR6	15690042	selective and non-redundant sensor of microbial diacylglycerides. TLR6-CD36-TLR4 activation is a common molecular mashanime hu which athenoamic linida and amulaid
13675	TLR6	20037584	mechanism by which atherogenic lipids and amyloid- beta stimulate sterile inflammation. TLR2::TLR6 synergistically interacts with TLR9 in lung epithelium to induce rapid pathogen killing, and
13675	TLR6	21482737	can be used as a therapeutic target to treat otherwise lethal pneumonia. Cutaneous bacteria can negatively regulate skin-driven immune responses by inducing Gr1(+)CD11b(+)
13675	TLR6	25456159	<ul><li>myeloid-derived suppressor cells via TLR2-6 activation.</li><li>Cxcl16-Cxcr6 crosstalk coordinates the intestinal topography of Il22 secretion required for mucosal</li></ul>
206490	Cxcr6	25456160	defence against Citrobacter rodentium infection. Cxcl16 is a chemoattractant in cerebrospinal fluid in early pneumococcal meningitis where Cxcl16 was found to be upregulated in RAW264.7 macrophages (but not in neutrophils and endothelial cells) upon pneumococcal stimulation. Cxcl16 upregulation in
194380	Cxcl16	20874518	vivo was dependent on Toll-like receptor (TLR) 2/TLR4 and MyD88 signaling. Cxcl16-Cxcr6 crosstalk coordinates the intestinal
194380	Cxcl16	25456160	topography of Il22 secretion required for mucosal defence against Citrobacter rodentium infection. DEFA6 (HD-6) and DEFA5 (HD-5) are two Paneth cell alpha-defensins found in the gut and alpha-
6012	DEFA6	19024344	defensing have multiple functions in the immune system. DEFA6 is secreted by paneth cells in the small intestine and is secreted in response to cholinergic and microbial stimuli. DEFA5A confer immunity to oral infection by Salmonella and is a major determinant of the small integrinal microbiome composition
6012	DEFA6	21560070	the small intestinal microbiome composition.

			DEFA6 is involved in mucosal innate immunity by protecting the small intestine against invasion by diverse enteric pathogens. In response to bacterial surface proteins, DEFA6 undergoes ordered self- assembly to form fibrils and nanonets that surround
6012	DEFA6	22722251	and entangle bacteria. NOD2 plays a role in intestinal innate immunity by regulating the expression of DEFA5 and DEFA6
6012	DEFA6	25433720	through the NF-kB and MAPK pathways. DEFA5 and DEFA6 are two Paneth cell alpha- defensins found in the gut that have multiple functions
6199	DEFA5	19024344	in the immune system. DEFA5 expression in mice showed significant loss of segmented filamentous bacteria, thus demonstrating a novel role for Paneth cell defensins in intestinal homeostasis by regulating the small intestinal
6199	DEFA5	21468224	microbiome. DEFA5 is secreted by paneth cells in the small intestine and is secreted in response to cholinergic and microbial stimuli. DEFA5A confer immunity to oral infection by Salmonella and is a major determinant of the small intestinal microbiome composition.
6199	DEFA5	21560070	NOD2 plays a role in intestinal innate immunity by regulating the expression of DEFA5 and DEFA6
6199	DEFA5	25433720	through the NF-kB and MAPK pathways. NOD2 and NOD1 represent central players in the
30654	NOD2	17690884	control of the immune responses to bacterial infections and inflammation. NOD2 is an intracellular receptor of muramyl dipeptide (MDP), a component of peptidoglycan present in the cell wall of Gram-positive (G+) and Gram-negative
30654	NOD2	18240302	(G-) bacteria. NOD2 and NOD1 can induce CCL5 (RANTES) through NF-kappaB pathway, orchestrating the global Nod-dependent immune defence during bacterial
30654	NOD2	17705131	infections. NOD2-dependent recognition of S. aureus and muramyl dipeptide is facilitated by alpha-toxin (alpha- hemolysin), a pore-forming toxin and virulence factor of the pathogen and is dependent on IL-1beta-amplified
30654	NOD2	19541630	production of IL-6. NOD2 and NOD1 direct autophagy by recruiting ATG16L1 to the plasma membrane at the site of
30654	NOD2	19898471	bacterial entry. NOD2 stimulation induces autophagy in dendritic cells
30654	NOD2	19966812	influencing bacterial handling and antigen presentation. NOD2 recruits the critical autophagy protein ATG16L1
30654	NOD2	20200479	to the plasma membrane during bacterial invasion.

30654	NOD2	19701189	NOD2 functions as a cytoplasmic viral pattern- recognition receptor (PRR) and activates an innate immune responses to viral ssRNA by triggering activation of interferon-regulatory factor 3 (IRF3) and production of interferon-beta (IFNB1). NOD2 is both a positive and negative regulator of TLR4 - the effect it exerts is dependent on the presence of MDP. NOD2 upon engagement with its ligand, MDP, positively regulates TLR4-mediated signalling; in the absence of MDP, NOD2 negatively regulates the TLR4 pathway. (Demonstrated in murine model)
30654	NOD2	21199260	NOD2 is a peripheral peptidoglycan intracellular sensor and is important for the progression and pathogenesis of experimental autoimmune encephalomyelitis (animal model of multiple sclerosis).
30654	NOD2	21236705	NOD2 detects heat-killed Legionella pneumophila and stimulates NFkB and IFN-beta promoter activity. NOD2 deficiency results in increased proinflammatory cytokine expression at 4hrs and greater neutrophil recruitment to the lung. (Demonstrated in murine model)
30654	NOD2	21108472	DDX58 and NOD2 colocalize to cellular ruffles and cell-cell junctions to form a protein complex via the CARD domains. DDX58 negatively regulates ligand- induced NFkB signalling mediated by NOD2, and conversely, NOD2 negatively regulates type I
30654	NOD2	21690088	interferon induction by DDX58. NOD2 recognition of muramyl dipeptide, a component of bacterial cell walls, improves the barrier function of
30654	NOD2	22750073	intestinal epithelial cells. (Demonstrated in mice) NOD2 enhances the innate immune response of alveolar macrophages to Mycobacterium tuberculosis
30654	NOD2	22531915	in human. Following NOD2 activation, IRF4 interacts with MYD88, TRAF6, and RIPK2 and downregulates K63- linked polyubiquitinylation of RICK and TRAF6
30654	NOD2	24670424	leading to disruption of NFkB activation pathways. NOD2 plays a role in intestinal innate immunity by regulating the expression of DEFA5 and DEFA6
30654	NOD2	25433720	through the NF-kB and MAPK pathways. Adam17 affects sensitivity to interleukin-1 by
148186	Il1r2	25461404	changing the balance between Illr1 and Illr2 receptors. Illr1 is the primary receptor for the inflammatory
148254	Il1r1	22426547	cytokine II1b. (Demonstrated in human) IL1R1 upregulates Mir135b as a negative feedback regulatory mechanism to resolve cigarette smoke-
148254	Il1r1	23440414	induced inflammation in the lung.

			In response to adenovirus infection, the IL1A-IL1R1- CXCR2 signalling axis cooperates with complement to recruit Ly-6G+7/4+ polymorphonuclear leukocytes to the splenic marginal zone (MZ) in the proximity of
148254	Il1r1	24651866	virus-containing MARCO+ residential MZ macrophages, which are subsequently eliminated.
148254	Il1r1	25461404	Adam17 affects sensitivity to interleukin-1 by changing the balance between Il1r1 and Il1r2 receptors. Adam17 affects sensitivity to interleukin-1 by
130508	Adam17	25461404	Adam17 affects sensitivity to interleukin-1 by changing the balance between II1r1 and II1r2 receptors. IRF1 is required for the up-regulation of the CD40-NF-kappaB activator 1 (TRAF3IP2) axis during airway
42125	IRF1	12759449	inflammation. IRF1 is controlled by two distinct signalling pathways; a JAK/STAT-signalling pathway in viral infected cells
42125	IRF1	12420214	and an ATM-signalling pathway in DNA damaged cells. IRF1 specifically binds to the upstream regulatory
			region of the human IFN-beta (IFNB1) gene and mediates IFNB1 virus-induced transcription and is involved in the regulation of other genes such as IFN-
42125	IRF1	3409321	alpha and MHC class I genes. IRF1 mediated type I IFN independent mechanism of enhanced RSAD2 (viperin) expression provides a
42125	IRF1	20308629	redundant mechanism to protect cells from viral infections. IRF1 transcriptionally inhibits the IL23A through the ISRE element and reduce the severity of chronic intestinal inflammation caused by LPS (shown in
42125	IRF1	21097874	mice).
42125	IRF1	22266972	IRF1 promotes immune cell apoptosis and inhibits autophagy in a murine endotoxemia model. (Demonstrated in mice) MIR23A contributes to human herpes simplex virus
42125	IRF1	25461762	<ul><li>type 1 replication through the regulation of the IRF1- mediated antiviral signal pathway.</li><li>MIR23A contributes to human herpes simplex virus</li></ul>
126931	MIR23A	25461762	type 1 replication through the regulation of the IRF1- mediated antiviral signal pathway.
171492	1115	22084435	<ul><li>II15 regulates homeostasis and terminal maturation of NKT cells.</li><li>IL15 secreted by inflammatory monocytes is critical for</li></ul>
171492	1115	22940097	the differentiation of memory CD8(+) T and NK lymphocytes into antimicrobial effector cells. Cxcr3 expression in innate CD8+ T cells defines
171492	1115	25466888	protective antibacterial and cancer immunity upon Il15 stimulation. Il28ra (Ifnlr1), Stat1 and Irf3 are required for
197156	Il28ra	25431490	antibiotics to prevent persistent murine norovirus infection.

			The Nlrc4 inflammasome is important for control of mucosal Candida infection, impacting on inflammatory cell recruitment to infected tissues, as well as
197890	Nlrc4	22174673	protecting against the systemic dissemination of infection. Flagellin-induced Nlrc4 inflammasome activation in
197890	Nlrc4	22231517	splenic dendritic cells triggers antigen-independent IFN-gamma production by memory T cells. Nlrc4 is important for host survival and bacterial clearance, as well as neutrophil-mediated inflammation
197890	Nlrc4	22547706	in the lungs following Klebsiella pneumoniae infection. Nlrc4-dependent production of Il1b by intestinal phagocytes is a mechanism that discriminates pathogenic from commensal bacteria in the intestinal
197890	Nlrc4	22484733	host defence. NIrc4 Ser â € 533 phosphorylation is essential for
197890	Nlrc4	22885697	procaspase-1 recruitment to the Nlrc4 inflammasome complex after S. typhimurium infection. Microglial cells employ the NAIP5/NLRC4
197890	Nlrc4	23355222	inflammasome to monitor and clear central nervous system infections by flagellated bacteria. Flagellin induces Tlr5-dependent Il22 production and Nlrc4-dependent Il18 production to promote a protective gene expression program in intestinal
197890	Nlrc4	25395539	epithelial cells and elimination of rotavirus-infected cells. Actin polymerization is required for Nlrc4-dependent regulation of intracellular bacterial burden, inflammasome assembly, pyroptosis, and Illb
197890	Nlrc4	25422455	production. Nod1 and Nod2 activation results in substantial secretion of Ccl5 by murine macrophages and induces
149298	Nod1	17705131	binding of NF-kappaB subunits to Ccl5 promoter. Nod1 can activate the ISGF3 signaling pathway that is usually associated with protection against viral infection to provide mice with robust type I IFN- mediated protection from H. pylori and possibly other
149298	Nod1	20389019	mucosal infections. Nod1 and Nod2 account for neutrophil recruitment to
140202	No.41	20605241	the lungs of mice infected with Legionella pneumophila.
149298	Nod1	20685341	Nod1 and Nod2 can detect Legionella pneumophila and these receptors modulate the in vivo pulmonary
149298	Nod1	21072876	immune response differently.

149298	Nod1	21236705	Nod1 is a peripheral peptidoglycan intracellular sensor and is important for the progression and pathogenesis of experimental autoimmune encephalomyelitis (animal model of multiple sclerosis).
149298	Noui	21230703	Nod1 detects heat-killed Legionella pneumophila and stimulates NFkB and IFN-beta promoter activity. Nod1 deficiency results in impaired bacterial clearance and increased proinflammatory cytokine at 24hrs post-
149298	Nod1	21108472	infection. Nod1 is expressed by trophoblast cells across gestation and may have a role in mediating infection-associated inflammation and prematurity. Activation of Nod1 by
149298	Nod1	21677137	bacterial peptidoglycan-derived peptide induces maternal-fetal inflammation and preterm labour. Nod1 KO mice were protected from high-fat diet induced inflammation, lipid accumulation, and peripheral insulin intolerance. Ex vivo, Nod1 activation by bacterial peptidoglycan mimetics induces proinflammatory cytokine secretion and impaired insulin-stimulated glucose uptake in adipocytes. Hence, Nod1 is a plausible, new link between innate immunity
149298	Nod1	21715553	and metabolism. Nod1 and Nod2 synergize with Tlr4 in dendritic cells to increase IL12 production and enhance invariant natural killer T (iNKT) cell activation, and are important regulators of the IFN gamma response by iNKT cells during S. typhimurium and L.
149298	Nod1	24163408	monocytogenes infections. Salmonella enterica serovar Typhimurium Î mmsbB that possesses a modified lipid A triggers exacerbated colitis in the absence of Nod1 and/or Nod2, which is
149298	Nod1	25423082	likely due to increased Tlr2 stimulation. TGF-Î <sup>2</sup> signalling upregulates MIR181A2 expression
3485	SMAD4	25410655	through SMAD3/4-dependent promoter activation. TGF- $\hat{1}^2$ signalling upregulates MIR181A2 expression
18366	SMAD3	25410655	through SMAD3/4-dependent promoter activation. TGF- $\hat{1}^2$ signalling upregulates MIR181A2 expression
126715	MIR181A2	25410655	through SMAD3/4-dependent promoter activation. LCN2 is secreted by the urinary tract mucosa in response to uropathogenic E. coli challenge and acts in
87415	LCN2	25398327	innate immune defences as a colonization barrier. Tyrobp impairs host defence during pneumococcal
168769	Tyrobp	25402298	pneumonia at the primary site of infection by inhibiting phagocytosis by alveolar macrophages. Relb is required for II17a production in T cell in response to bacterial infection. Relb deficient T cells
151419	Relb	21419662	resulted in a diminished innate immune response to E. coli infection. Ripk3 has a novel function in NF-kB activation,
151419	Relb	25367573	dendritic cell biology, innate inflammatory-cytokine expression, and injury-induced tissue repair.

			Nfkb1 enforces specificity of cellular response to pathogens by binding to a subset of IRE sequences in IFN-inducible genes. Nfkb1 deficiency results in the
195445	Nfkb1	21343618	inappropriate production of Ifnb in response to bacterial DNA sensed by Tlr9. Mir126-Kdr axis is an important regulator of the innate response. Mir126 controls the survival and function of
195445	Nfkb1	24270517	plasmacytoid dendritic cells and regulates gene expression of Tlr7, Tlr9, Nfkb1 and Kdr. Ripk3 has a novel function in NF-kB activation, dendritic cell biology, innate inflammatory-cytokine
195445	Nfkb1	25367573	expression, and injury-induced tissue repair. Ripk3 forms a complex with Ticam1 upon Toll-like receptors (TLR) 3 and 4 activation resulting in Ripk3- dependent but TNF-independent necrosis in
165974	Ripk3	22123964	macrophages. Ripk3 interacts with Zbp1 to mediate virus-induced
165974	Ripk3	22423968	necrosis. Ripk3 has a novel function in NF-kB activation, dendritic cell biology, innate inflammatory-cytokine
165974	Ripk3	25367573	expression, and injury-induced tissue repair. SCN5A is a novel pathogen sensor that initiates anti- viral signalling and transcription through ADCY8 and ATF2.
75609	ATF2	25368329	SCN5A is a novel pathogen sensor that initiates anti- viral signalling and transcription through ADCY8 and ATF2.
35780	ADCY8	25368329	SCN5A is a novel pathogen sensor that initiates anti- viral signalling and transcription through ADCY8 and ATF2.
26059	SCN5A	25368329	
29564	ECSIT	10465784	ECSIT is specific for the Toll/IL-1 pathways and is a regulator of MAP3K1 (MEKK1) processing. ECSIT binds to MAP3K7 and TRAF6 to form a
29564	ECSIT	25371197	complex that plays a pivotal role in activating TLR4- mediated NF-kB signalling.
166877	Slc11a1	25350459	Slc11a1 confers temporal and anatomical host resistance to chronic Salmonella infection. USP2 deubiquitinates K63-linked polyubiquitin chains from TBK1 to terminate TBK1 activation and negatively regulate IFNB1 signalling and antiviral
74471	USP2	25070846	<ul><li>Pituitary hormone prolactin (Prl) constrains tumor- promoting liver inflammation by inhibiting Map3k1- dependent activation of Myc at the level of the</li></ul>
139889	Prl	25049387	trafasome.

			Pituitary hormone prolactin (Prl) constrains tumor- promoting liver inflammation by inhibiting Map3k1- dependent activation of Myc at the level of the trafasome.
144556	Мус	25049387	ARG1 expression induced by intracellular pathogens in mouse classically activated macrophages (CAMs) through the TLR pathway, suppressing nitric oxide
96496	ARG1	18978793	production and thwarting effective immunity independent of the STAT6 pathway. PI3K/PTEN-regulated extracellular ARG1 acts as a paracrine regulator of inflammation and immunity. Demonstrated in mice.
96496	ARG1	25015834	PI3K/PTEN-regulated extracellular ARG1 acts as a paracrine regulator of inflammation and immunity. Demonstrated in mice.
81409	PTEN	25015834	Demonstrated in mee.
133881	Pik3cg	25022365	Rab8a interacts with Pik3cg to regulate Akt signalling generated by surface Tlr4. Rab8a interacts with Pik3cg to regulate Akt signalling
167493	Rab8a	25022365	generated by surface Tlr4. STAT1 mediates IFNG cytokine signalling, and as a member of the STAT family protein, STAT1 has a significant impact on the innate immunity during
77617	STAT1	17971840	sepsis. STAT1 serine phosphorylation is induced by TNF- alpha and PGE2 and this activates expression of the
77617	STAT1	18678606	STAT1 and NF-kappaB target gene IFN regulatory factor 1 (IRF1), which contributes to IFN responses. STAT1 phosphorylation and DNA binding activity is under the control of Janus protein-tyrosine kinases
77617	STAT1	7657660	(JAKs) and the epidermal growth factor receptor (EGFR). Acetylation of STAT1 modulates NF-kappaB activity
77617	STAT1	16481475	and thus ultimately apoptosis where, as a result, RELA DNA binding, nuclear localization, and expression of anti-apoptotic NF-kappaB target genes decrease. STAT1 is part of the interferon-stimulated gene factor 3 (ISGF3) transcription complex which is composed of
77617	STAT1	8943351	a STAT1:2 heterodimer and a weak DNA-binding protein, IRF9. STAT1 transcription factor enhances TLR8
77617	STAT1	20829351	functionality by binding of to GAS elements on the TLR8 promoter in an IFN-gamma-dependent manner. STAT1 phosphorylation at Ser708 is a key event in the IFN signalling pathway that imparts anti-viral
77617	STAT1	22065572	immunity to restrict West Nile virus infection. (Demonstrated in mice)
77617	STAT1	22425562	Histone deacetylase inhibitors prevent IFNG-mediated phosphorylation of STAT1.

			TNK1 is a component of the IFN-JAK-STAT signalling cascade and is a critical antiviral host factor where its abundance is inversely correlated to viral replication and contributes to the hepatocytic response
77617	STAT1	24449862	to antiviral treatment. STAT1 is directly recruited to TRAF6, demonstrating cross-talk between the TLR and JAK/STAT signalling pathways, and this direct activation of STAT1 by TLR
77617	STAT1	25027037	signalling suggests a crucial role for STAT1 in TLR- induced inflammation. Demonstrated in mice. MAP3K7 (TAK1) Ser412 phosphorylation is regulated by PRKACA and PRKX, and is essential for proper
41523	PRKX	25028512	signalling, as well as proinflammatory cytokine induction by TLR/IL-1R activation. MAP3K7 (TAK1) Ser412 phosphorylation is regulated by PRKACA and PRKX, and is essential for proper
32929	PRKACA	25028512	signalling, as well as proinflammatory cytokine induction by TLR/IL-1R activation. T-cell-intrinsic Mir155 is required for type-2 immunity, in part through regulation of S1pr1, whereas
189693	S1pr1	25024218	T-cell-intrinsic Mir146 is required to prevent overt Th1/Th17 skewing. CXCL2 is a chemokine secreted by kidney dendritic
24257	CXCL2	21757770	cells to recruit neutrophils to sites of uropathogenic bacterial infection. (Demonstrated in mouse) Secreted CCNA2 (CCN1) promotes anti-inflammatory cytokine IL10 release from epithelial cells via integrin $\hat{i} + N\hat{i}^2$ ( DKC) and this subsequently supersess TNF
24257	CXCL2	25005359	$\hat{l}\pm V\hat{l}^2$ 6-PKC, and this subsequently suppresses TNF, CXCL2 and neutrophil infiltration in the lungs. Secreted CCNA2 (CCN1) promotes anti-inflammatory cytokine IL10 release from epithelial cells via integrin $\hat{l}\pm V\hat{l}^2$ 6-PKC, and this subsequently suppresses TNF,
36372	CCNA2	25005359	CXCL2 and neutrophil infiltration in the lungs. MIR124-1 plays a negative regulatory role in fine- tuning the inflammatory response in alveolar macrophages upon mycobacterial infection, in part by
126971	MIR124-1	24995397	directly targetting TLR signalling. PIK3CA, as the alpha subunit of PI3K, functions as a negative regulator of TLR signalling. The activation of
66003	PIK3CA	17827709	PIK3CA results in the inhibition of pro-inflammatory events such as expression of IL12 and TNFA. PIK3CA and PIK3CB isoforms of class IA
66003	PIK3CA	20953381	phosphatidylinositol 3-kinase (PI3K) are both required for the pro-inflammatory response to flagellin.
66003	РІКЗСА	24996183	MIR203A accelerates apoptosis in LPS-stimulated alveolar epithelial cells by targeting PIK3CA. MIR203A post-transcriptionally regulate the pro- information and H 24 in
126601	MIR203	22917968	inflammatory cytokines TNFalpha and IL24 in keratinocytes.
126601	MIR203	23785202	MIR203A can be induced by IFN in virally infected cells to dampen IFN anti-viral signalling.

126601	MIR203	24996183	MIR203A accelerates apoptosis in LPS-stimulated alveolar epithelial cells by targeting PIK3CA. Trem1 expression is upregulated following Pseudomonas aeruginosa infection in the cornea and the inhibition of Trem1 reduces the severity of corneal disease. Trem1 acts as an inflammatory amplifier in P. aeruginosa keratitis by modulating TLR signalling and Th1/Th2 responses.
189794	Trem 1	21555403	Trem1 expression is induced by vitamin D3 in human bronchial epithelial cells. Activation of TREM1 leads to the induction of human beta defensin 2 and TNF- alpha mRNA in the airway epithelium. (Demonstrated
189794	Trem1	21690199	in human) Trem1 mediates endotoxin tolerance in monocytes through its ability to induce anti- or pro-inflammatory signals depending on its membrane-bound state.
189794	Trem1	22459945	(Demonstrated in human) Trem1 offers protective innate immunity during pneumococcal pneumonia by enhancing the early
189794	Trem1	24752755	immune response of alveolar macrophages. SAMHD1 is specifically targeted for degradation by human immunodeficiency virus 1 (HIV-1) viral protein VPX, and mutations in SAMHD1 causes Aicardi- Goutieres syndrome, which mimics symptoms of
73653	SAMHD1	21720370	congenital viral infection. SAMHD1 is expressed in dendritic and myeloid cells and acts as an anti-retroviral protein that inhibits the early stages of the viral life cycle. Silencing of SAMHD1 leads to significant accumulation of human immunodeficiency virus 1 (HIV-1) DNA in infected
73653	SAMHD1	21613998	cells. SAMHD1 inhibits HIV replication by depleting intracellular dNTPs, which are required for viral
73653	SAMHD1	22327569	reverse transcriptase to synthesize viral DNA. Full activation of SAMHD1 involves ordered binding of GTP and substrate dNTPs to activator and substrate
73653	SAMHD1	24753578	sites on the enzyme. In response to adenovirus infection, the IL1A-IL1R1- CXCR2 signalling axis cooperates with complement to recruit Ly-6G+7/4+ polymorphonuclear leukocytes to the splenic marginal zone (MZ) in the proximity of virus-containing MARCO+ residential MZ
166395	Cxcr2	24651866	macrophages, which are subsequently eliminated. Fibrates protect against bacteria†induced sepsis by inhibiting LPS†mediated ERK phosphorylation and the expression of Adrbk1 (GRK2), thereby preventing
166395	Cxcr2	24755316	<ul> <li>Cxcr2 down-regulation on neutrophils.</li> <li>Fibrates protect against bacteria†induced sepsis by inhibiting LPS†mediated ERK phosphorylation and the expression of Adrbk1 (GRK2), thereby preventing</li> </ul>
129419	Adrbk1	24755316	Cxcr2 down-regulation on neutrophils.

126375	MIR122	23580661	Circulating MIR122 activates natural killer cells via the TLR1 signalling pathway. Reduction of MIR122 expression may play a role in the
126375	MIR122	24672032	progression of fibrosis in patients with chronic hepatitis C.
127039	hsa-mir-146a	19840932	MIR146A is critical for in vitro monocytic cell-based endotoxin tolerance. MIR146A is involved in control of Toll-like receptor and cytokine signaling through a negative feedback regulation loop involving down-regulation of IL-1 receptor-associated kinase 1 (IRAK1) and TNF
127039	hsa-mir-146a	16885212	receptor-associated factor 6 (TRAF6) protein levels. MIR146A-mediated and NF-kappaB-sensitive inflammatory circuit was found in Alzheimer disease and in stressed human brain cells.
127039	hsa-mir-146a	18801740	
			MIR146A expression is driven by the transcription factor NF-kappaB and changes in the expression of hsa-mir-146a have been implicated in both the development of multiple cancers and in the negative regulation of inflammation induced via the innate
127039	hsa-mir-146a	19021527	immune response. MIR146A negatively regulates the IL-1-beta-induced
127039	hsa-mir-146a	19021527	release of chemokines IL-8 and RANTES. MIR146A negatively regulates the type I IFN pathway
127039	hsa-mir-146a	19333922	by directly targeting IRF5 and STAT1. MIR146A feedback inhibits RIG-I-dependent type I
127039	hsa-mir-146a	19596990	IFN production in macrophages by targeting TRAF6, IRAK1, and IRAK2. MIR146A significantly increases after 24 h stimulation in monocytes from neonate cord blood compared to those derived from adults, suggesting the involvement
127039	hsa-mir-146a	20616571	of microRNA in immune regulation of the innate immune system of neonates. MIR146A-mediated down-regulation of IRAK-1 coupled to an NF-ΰB-induced up-regulation of IRAK-2
127039	hsa-mir-146a	20937840	expression drives an extensively sustained inflammatory response. MIR146A mediates the intestinal epithelial innate immune tolerance during the neonatal period, characterizing tolerance as an active condition involved
127039	hsa-mir-146a	20951969	characterizing tolerance as an active condition involved in the establishment of intestinal mucosal homeostasis. MIR146A negatively regulates both oxidized low- density lipoprotein accumulation and the inflammatory
127039	hsa-mir-146a	21329689	response in macrophages by inhibiting TLR4 and the activation of TLR4-dependent signalling pathways. MIR146A is highly expressed in patients with Sjogren's syndrome and was found to increase the phagocytic activity and suppressing the inflammatory
127039	hsa-mir-146a	21469088	cytokine production in human monocytes.

MIR146A transcription is induced by NFkB in response to stimulations such as LPS, TNF-alpha and IL1B. MIR146A targets TRAF6 and IRAK1 and act s as a negative regulator fine-tuning the immune response.

MIR146A directly targets RELB to modulate the amplitude of monocyte response to inflammatory 127039 hsa-mir-146a 22545247 challenges. MIR146A is a mechanosensitive miRNA that modulates mechanotransduction and pressure-induced 127039 hsa-mir-146a 22593544 inflammation in small airway epithelium. MIR146A is overexpressed during prion disease and modulates the innate immune response and the 127039 hsa-mir-146a 22363497 microglial activation state. (Demonstrated in mice) MIR146A is upregulated in lung cells in response to 127039 infection with influenza A virus. hsa-mir-146a 22822053 MIR146A is a key regulator of astrocyte-mediated 127039 hsa-mir-146a 23028621 inflammation responses in neurological disorders. MIR146A alleviates ischemia-induced inflammation in 127039 hsa-mir-146a 23143987 the small intestine by downregulating IRAK1. In the course of influenza A infection MIR146A 127039 hsa-mir-146a 23343627 transcription increases to limit viral propagation. MIR146A targets components of the NF-kB and EGR 127039 hsa-mir-146a 23733368 pathways to repress inflammation. MIR146A is a potent negative regulator of the innate keratinocytes immune response in through 127039 hsa-mir-146a 24670381 downregulation of the IRAK1/TRAF6/NFÎ<sup>o</sup>B pathway. RIPK2 functions in innate immunity by mediating NOD1 and NOD2 signalling but not TLR-mediated 27941 RIPK2 17277144 immune responses. RIPK2 polyubiquitination mediates the recruitment of MAP3K7 (TAK1) and the subsequent activation of NF-kappaB signalling via IKK complex in NOD 27941 RIPK2 18079694 signalling. RIPK2 deubiquitination by TNFAIP3 (A20) restricts 27941 RIPK2 18342009 NF-kappaB activation via NOD2 signalling. RIPK2 is sequestered by MAP3K4 to inhibit the NOD2:RIPK2 complex from activating NF-kappaB 27941 RIPK2 18775659 signalling pathways. RIPK2 is a downstream adaptor molecule in the NOD1/2 signaling pathway and is important for the and pathogenesis of experimental progression autoimmune encephalomyelitis (animal model of multiple sclerosis). RIPK2 was found to be critical for the activation of CNS-infiltrating dendritic cells. 27941 RIPK2 21236705 RIPK2 is essential for NOD1 and NOD2-signalling upon recognition of bacterial peptidoglycan. RIPK2 is crucial for inflammatory cytokine secretion, activation and recruitment of macrophage and neutrophils as well as the capacity to activate the adaptive immune

response. (Demonstrated in murine model)

127039

27941

RIPK2

hsa-mir-146a

21652514

21469090

			Following NOD2 activation, IRF4 interacts with MYD88, TRAF6, and RIPK2 and downregulates K63-
27941	RIPK2	24670424	linked polyubiquitinylation of RICK and TRAF6 leading to disruption of NFkB activation pathways. IRF4 participates in the regulation of lymphoid cell
55681	IRF4	12566414	apoptosis by modulating the efficiency of the Fas- mediated death signal. IRF4 provides a positive feedback signal for its own gene expression in dendritic cells (DCs) and the
55681	IRF4	16272311	expression of IRF4 mRNA, but not of other IRFs, is specifically up-regulated during DC differentiation. Following NOD2 activation, IRF4 interacts with MYD88, TRAF6, and RIPK2 and downregulates K63-
55681	IRF4	24670424	linked polyubiquitinylation of RICK and TRAF6 leading to disruption of NFkB activation pathways. Astrocytes produce IL19 and express IL19 receptors (Il20ra and Il20rb) in response to bacterial pathogens.
193159	Il20rb	24677051	IL19 can significantly attenuate bacterially induced inflammatory astrocyte responses Astrocytes produce IL19 and express IL19 receptors (Il20ra and Il20rb) in response to bacterial pathogens.
136067	Il20ra	24677051	IL19 can significantly attenuate bacterially induced inflammatory astrocyte responses Astrocytes produce IL19 and express IL19 receptors (Il20ra and Il20rb) in response to bacterial pathogens.
190663	Il19	24677051	IL19 can significantly attenuate bacterially induced inflammatory astrocyte responses Ptges (prostaglandin E2) is produced by LPS-primed macrophages upon treatment with silica crystal and
157032	Ptges	21497116	aluminum salts, and is important for the production of IgE in Th2 cells. Influenza A virus infection targets the Ptges and
157032	Ptges	24726877	prostaglandin E2 pathway to evade host type I IFN- dependent antiviral immunity Hepatitis C infection stimulates the expression of DNM1L and MFF and promotes DNM1L recruitment
82567	MFF	24733894	to mitochondria by stimulating the phosphorylation of DNM1L, leading to mitochondrial fission. Hepatitis C infection stimulates the expression of DNM1L and MFF and promotes DNM1L recruitment
26348	DNM1L	24733894	to mitochondria by stimulating the phosphorylation of DNM1L, leading to mitochondrial fission. BST2 directly binds to purified LILRA4 (ILT7) protein, initiates signalling via the ILT7-
36619	BST2	19564354	FcepsilonRIgamma complex, and strongly inhibits production of IFN and pro-inflammatory cytokines in plamsmacytoid dendritic cells BST2 (CD317) is a broadly acting and conserved mediator of innate control of retroviral infection and pathogenesis that restricts the release of retroviruses
36619	BST2	20702620	and lentiviruses in rodents by limiting replication of HIV-1 and MLV.

36619	BST2	21734563	BST2 is an innate intracellular HIV restriction factor that is upregulated by type I interferons. BST2 is a broad spectrum effector of the innate immune response to viral infection. BST2 is antagonized by the human immunodeficiency virus
36619	BST2	22072710	<ul><li>(HIV) Vpu protein to evade innate immune system detection.</li><li>HIV-1 viral protein U, Vpu, protects HIV-infected cells</li></ul>
36619	BST2	24733916	from antibody-dependent cell-mediated cytotoxicity as a function of its ability to counteract BST2 (tetherin). Mir302b expression is up-regulated upon bacterial
221396	Mir302b	24717937	infection and is a crucial regulator of NFΰB signalling by directly targeting IRAK4 MAPK9 phosphorylates IRF3 and is essential for IRF3 dimerization induced by polyinosinic-cytidylic acid
62183	МАРК9	19153595	<ul><li>(polyI:C).</li><li>Upon viral infection, MAVS recruits MKK7 onto mitochondria, leading to the induction of apoptosis by</li></ul>
62183	МАРК9	24651600	MAP2K7 activated MAPK9 Upon viral infection, MAVS recruits MKK7 onto
24000	MAP2K7	24651600	mitochondria, leading to the induction of apoptosis by MAP2K7 activated MAPK9 Chitin induces IL25, IL33, and TSLP which are required to stimulate ILC2 production of IL5 and IL13. IL5 and IL13, in turn, are required for the accumulation
172199	115	24631157	of eosinophils and alternatively activated macrophages that are associated with allergy. Il13 dampens the innate immune response in airway epithelial cells via Irak3-mediated inhibition of Tlr2
172064	II13	22154382	signalling. (Demonstrated in human) IL13 produced by activated innate lymphoid cells type 2 (ILC2s) is critical for promoting the migration of activated lung dendritic cells to the draining lymph node, and in the differentiation of naive CD4+ T cells
172064	II13	24613091	into Th2 cells The noncanonical NF ΰB pathway regulates histone modifications at the Ifnb1 promoter resulting in attenuated recruitment of Rela and histone demethylase, Kdm4a, to the Ifnb1 promoter. This provides a mechanism for regulating the induction of
180637	Kdm4a	24656046	type I interferons . Flt4 signalling represents a negative feedback mechanism by immune cells to overreacting during bacterial infection; ligation of Flt4 by Vegfc reduces
155240	Vegfc	24656836	TLR4-triggered production of proinflammatory cytokines in macrophages. Flt4 signalling represents a negative feedback mechanism by immune cells to overreacting during bacterial infection; ligation of Flt4 by Vegfc reduces
167138	Flt4	24656836	TLR4-triggered production of proinflammatory cytokines in macrophages.

			FLT4 signalling represents a negative feedback mechanism by immune cells to overreacting during bacterial infection; ligation of FLT4 by VEGFC reduces TLR4-triggered production of proinflammatory
45463	VEGFC	24656836	cytokines in macrophages. FLT4 signalling represents a negative feedback mechanism by immune cells to overreacting during bacterial infection; where ligation of FLT4 by VEGFC
62355	FLT4	24656836	reduces TLR4-triggered production of proinflammatory cytokines in macrophages. Both Nlrp3 and Nlrp1a are important regulators of Toxoplasma proliferation and IL18 signaling is required to mediate host resistance to acute
195914	Nlrp1a	24549849	toxoplasmosis.
191058	Hdac1	24550390	Daxx interacts with Hdac1 and represses the transcription of II6 in TLR-triggered macrophages.
168138	Daxx	24550390	Daxx interacts with Hdac1 and represses the transcription of II6 in TLR-triggered macrophages. Il12, consisting of Il12a and Il12b subunits, induces Il2ra to form high-affinity Il2 receptors on natural killer
164545	Il12b	22888135	cells in response to mouse cytomegalovirus infection. Nod1 and Nod2 synergize with Tlr4 in dendritic cells to increase IL12 production and enhance invariant natural killer T (iNKT) cell activation, and are
164545	II12b	24163408	<ul><li>important regulators of the IFN gamma response by</li><li>iNKT cells during S. typhimurium and L.</li><li>monocytogenes infections.</li><li>CXCL10 exert direct antimicrobial effects in vitro</li><li>against Bacillus anthracis spore and bacilli in a</li><li>receptor-independent manner and contributes to</li><li>pulmonary innate immunity. (Demonstrated in murine</li><li>model)</li></ul>
25212	CXCL10	21124994	CXCL10 concentration in blood increases during neonatal polymicrobial sepsis, and the blockade of CXCL10 not only worsens recruitment and phagocytic function of macrophages, but also the survival of neonatal mice. (Demonstrated in murine model)
25212	CXCL10	21518789	MIR21 inhibition enhances CCL5 (RANTES) and CXCL10 (IP-10) release in MCF-7 cancer cells and
25212	CXCL10	23998932	resulted in increased lymphocyte migration. PIAS3 is a target of MIR21 in MCF-7 cells. ISG15 does not directly alter human rhinovirus replication but modulates immune signalling via the viral sensor protein DDX58 to impact production of
25212	CXCL10	24448099	<ul><li>CXCL10, which has been linked to innate immunity to viruses.</li><li>Human rhinovirus infection of epithelial cells induces the expression and secretion of ISG15, which modulates immune responses via effects on DDX58,</li></ul>
25212	CXCL10	24448099	and by regulating CXCL10 production.

148617	Prdm1	24477914	Prdm1 (Blimp1) modulates host defences by suppressing Ccl8-induced inflammation. The calpain-MYH9-RAB7B axis regulates TLR4
5777	МҮН9	24489676	containing alpha-granules trafficking in thrombin- stimulated platelets. THRIL, a large intergenic noncoding RNA, binds to heterogenous nuclear ribonucleoprotein L (HNRNPL) and forms a functional THRIL-HNRNPL complex that
49631	HNRNPL	24371310	regulates transcription of TNF by binding to its promoter. IFIT1 expression activates the IFN response during C. pneumoniae infection, mediated by intracellular nucleotide-sensing pattern recognition receptors
81960	IFIT1	20386592	(PRRs), which operate through a mechanism dependent on the bacterial type III secretion system (T3SS). IFIT1 recognizes and directly binds to 5'-triphosphate RNA and functions as an innate inhibitor of viral
81960	IFIT1	21642987	replication and pathogenicity. (Demonstrated in murine model) IFIT1 is an innate immune network bottleneck with the ability to suppress induction of TLR4 response genes
81960	IFIT1	22745654	<ul> <li>in LPS-stimulated macrophages. (Demonstrated in mice)</li> <li>IFIT1 binds mRNAs that lack 2â€<sup>2</sup>O methylation on the first ribose, thus inhibiting translation by impairing</li> </ul>
81960	IFIT1	24098121	binding of eukaryotic translation initiation factors to 2â€ <sup>2</sup> O-unmethylated RNA templates IFIT1 binds with high affinity to the cap-proximal regions of cap0-mRNAs abrogating 48S complex
81960	IFIT1	24371270	formation in an in vitro reconstituted translation system. BTK is a Toll/interleukin-1 receptor domain-binding
79651	ВТК	12724322	protein that participates in NF-kappaB activation by TLR4. BTK interacts with intracellular MHC class II molecules to activate adaptor molecules MYD88 and TICAM1 to promote TLR signalling. (Demonstrated in
79651	BTK	21441935	murine model) BTK is a positive regulator in the ITAM-mediated TREM1/TYROBP pathway, which induces pro- inflammatory cytokines such as TNF-alpha, IL8, and activation/differentiation cell surface markers. Patients suffering from X-linked agammaglobulinemia (XLA), which is a rare hereditary disease caused by mutation in the BTK gene, show reduced TNF-alpha induction in
79651	BTK	21659545	PBMCs upon TREM1 engagement. BTK directly phosphorylates TLR3 and plays a critical role in the induction of inflammatory cytokines and
79651	BTK	22454496	IFNB. (Demonstrated in mice) BTK is required for the activation of natural killer
79651	BTK	22589540	cells. (Demonstrated in mice)

			BTK is a negative regulator of TLR- or TNF- stimulated reactive oxygen species (ROS) production
79651	BTK	22366891	in neutrophils. BTK regulates TLR9-mediated induction of cytokines
79651	BTK	24375473	in plasmacytoid dendritic cells, but has no role in TLR7 signalling.
78146	TRIM14	23438823	TRIM14 was identified in a systematic screen for
/8140	1 K11114	23438823	positive regulators of innate immune responses. TRIM14 is a mediator of mitochondrial antiviral immunity facilitating the immune responses mediated by retinoic acid-inducible gene-I-like (RIG-I)-like
78146	TRIM14	24379373	receptors. Mir149 negatively regulates TLR/Myd88 mediated
221120	Mir149	24375488	inflammatory responses in macrophages by targeting Myd88 mRNA. Sykb promotes wound healing in human rhinovirus-
			infected airway epithelial cells. (Demonstrated in
152695	Sykb	22031919	human)
152695	Sykb	14699155	SYK is a tyrosine protein kinase that is an upstream activator of c-Jun N-terminal kinase (JNK). SYK interacts with PLCG2 and plays a role in
152695	Sykb	16449524	complement mediated phagocytosis by regulating both actin dynamics and the RhoA activation pathway. SYK interacts with TRAF-interacting protein (TIRAP),
152695	Sykb	19151749	two proteins with opposing effects on tumour necrosis factor (TNF) signalling where SYK enhances the activation of NF-kappaB by TNF and this is antagonized by TIRAP. SYK operating downstream of ITAM-coupled fungal pattern recognition receptors by controlling both pro-
152695	Sykb	19339971	IL-1beta synthesis and inflammasome activation after fungal infection. SYK and MYD88 adaptor protein pathways activation
152695	Sykb	19913447	by bacteria promotes regulatory properties of neutrophils. SYK is activated by several ITAM-containing or ITAM-coupled C-type lectin receptors on myeloid cells leading to the production of pro-inflammatory
152695	Sykb	20401526	cytokines including IL-1-beta to initiate antifungal responses. SYK promotes wound healing in human rhinovirus-
152695	Sykb	22031919	infected airway epithelial cells. Phosphorylation of the inflammasome adaptor Pycard (ASC) controls inflammasome activity through the formation of ASC specks. The NLRP3 and AIM2
152695	Sykb	24185614	inflammasomes require Syk and Mapk8 (JNK) for their full activity . The expression of Clec4e (Mincle) and its downstream
152695	Sykb	24212132	signal phospho-Syk/Syk increases after cerebral ischemia and reperfusion.

			Hyperactivated ERN1 (IRE1 $\hat{I} \pm$ ) increases TXNIP mRNA stability by reducing levels of a TXNIP destabilizing microRNA, miR-17. In turn, elevated
101710	TXNIP	22883233	<ul> <li>TXNIP protein activates the NLRP3 inflammasome, causing procaspase-1 cleavage and interleukin 11<sup>2</sup> (IL-11<sup>2</sup>) secretion.</li> <li>Endoplasmic reticulum (ER) stress-mediated reactive oxygen species accumulation leads to activation of</li> </ul>
101710	TXNIP	24217221	NLRP3 inflammasome through enhanced secretion of IL1B and binding of TXNIP. Krt16 participates in the regulation of early inflammation and innate immunity in a broad range of
211290	Krt16	24218583	settings involving skin, such as pachyonychia congenita and psoriasis. Cd51 (Aim) is required for obesity-associated
157061	Cd51	21730133	recruitment of inflammatory macrophages into adipose tissue. Cd51 contributes to the macrophage autophagy
157061	Cd51	24223991	mechanisms that lead to Mycobacterium. tuberculosis killing. The IFNL3 SNP rs4803217 is critical for the outcome
			of hepatitis C virus (HCV) infec¬tion by controlling the stability of IFNL3 mRNA. HCV induces two microRNAs, MIR499A and MIR208B, that target the polymorphic region of the IFNL3 3â€ <sup>2</sup> UTR.
273442	MIR208B	24241692	The IFNL3 SNP rs4803217 is critical for the outcome of
			hepatitis C virus (HCV) infec $\hat{A} \neg$ tion by controlling the stability of IFNL3 mRNA. HCV induces two microRNAs, MIR499A and MIR208B, that target the polymorphic region of the IFNL3 $3\hat{a} \in UTR$ .
126967	MIR499A	24241692	IFNL1, IFNL2 and IFNL3 (IL28B) have different
50088	IL28B	24041672	effects on Toll-like receptor-related gene expression in HepG2 cells. The IFNL3 SNP rs4803217 is critical for the outcome
			of hepatitis C virus (HCV) infec¬tion by controlling the stability of IFNL3 mRNA. HCV induces two microRNAs, MIR499A and MIR208B, that target the polymorphic region of the IFNL3 3â€ <sup>2</sup> UTR.
50088	IL28B	24241692	
207523	Fscn1	24244012	<ul> <li>Fscn1 contributes to the survival of dendritic cells against Listeria monocytogenes infection.</li> <li>Kaposi's sarcoma-associated herpesvirus (KSHV) infection results in upregulation of AICDA in primary human tonsillar B cells. Two KSHV miRNAs, K12-11 and K12-5 interact with the 3â€<sup>2</sup>UTR of AICDA to</li> </ul>
17253	AICDA	24244169	translationally repress its expression.

69007	PARD3	24244864	Atypical PKC and PARD3 are inhibitors of the canonical NF-ΰB activation pathway in epithelial cells. Lipid transfer proteins saposins play an essential role in modulating human invariant natural killer T cell reactivity to antigen-presenting cells activated by inflammatory stimuli. Lipid-loaded Saposin B mediates lipid transfer onto CD1D and accelerates
103733	CD1D	24248359	dissociation of CD1D-bound lipids, promoting lipid exchange. TICAM2 deficiency results in the impairment of LPS- stimulated TNF-alpha protein translation.
406966	TICAM2	21494017	(Demonstrated in murine model) ARF6 regulates LPS internalization and LPS-induced
406966	TICAM2	24297182	relocation of TICAM2 (TRAM), which is required for the MyD88-independent TLR4 signalling cascade. The homotypic interaction of TICAM2 Toll/interleukin-1 receptor (TIR) domain is essential to form a scaffold for recruiting the TICAM1 TIR
406966	TICAM2	24255114	domain. TICAM1 adaptor protein is displaced from TICAM2 by a splice variant of TICAM2, TAG, resulting in the
19505	TICAM1	19412184	negative regulation of the MyD88-independent TLR4 pathway. TICAM1 (TRIF)-dependent activation of CASP8 is involved in pro-apoptotic signalling through TLR3 and
19505	TICAM1	20019748	this under the control of inhibitor of apoptosis proteins in melanoma cells. TICAM1 preferentially activates the IFN-beta promoter
19505	TICAM1	12471095	in the Toll-like receptor signalling, particularly in the MyD88-independent pathway. TICAM1 (TRIF) recruits TRAF6-TAK1-TAB2 to TLR3 through its TRAF6-binding site, which is required for NF-kappaB but not IRF3 activation.
19505	TICAM1	14982987	TLR3/TICAM1-mediated NF-kappaB and IRF3 activation is induced by double-stranded RNA.
19505	TICAM1	12539043	TICAM-1 is an adaptor molecule that participates in TLR3-mediated interferon-beta induction. TICAM1 and TICAM2 both function in LPS-TLR4
19505	TICAM1	14517278	signalling to regulate the MyD88-independent pathway during the innate immune response to LPS. TICAM1 and TICAM2 form an adaptor complex that
19505	TICAM1	14519765	plays a crucial role in LPS-TLR4-mediated activation of IFN-beta. The TICAM1 signalling pathway in murine dendritic
19505	TICAM1	21454965	cells is crucial for dsRNA-mediated natural killer cell activation. (Demonstrated in murine model) TICAM1 deficiency results in the impairment of LPS-
19505	TICAM1	21494017	stimulated TNF-alpha protein translation. (Demonstrated in murine model)

TICAM1 is crucial for NLRP3 inflammasome activation in response specific to viable, but not heat-killed, E. coli infections. (Demonstrated in murine model)

19505	TICAM1	21602824	model
17505		21002021	TICAM1 is proteolytically cleaved by Enterovirus 71 3C to inhibit the induction of innate immunity by TLR3-signalling. TICAM1 cleavage results in the
19505	TICAM1	21697485	inhibition of NFkB and IFN-beta promoter activation. TICAM1 forms a dsRNA sensor complex with components DDX1, DDX21 and DHX36 to trigger the type I interferon and cytokine response to poly I:C, influenza A virus, and reovirus. (Demonstrated in
19505	TICAM1	21703541	murine model) TICAM1 is a potent negative regulator of TLR agonist- triggered immune responses, specifically suppressing IL12 in dendritic cells and IFNG in natural killer cells.
19505	TICAM1	21760953	(Demonstrated in mouse) TICAM1-TLR3-mediated signalling pathway plays an essential role in the anti-viral response against
19505	TICAM1	22072781	poliovirus infection. (Demonstrated in mouse) TICAM1 plays a role in host resistance to Gram- negative enteropathogens. TICAM1-mediated protective immunity is orchestrated by macrophage- induced IFN-beta and natural killer cell production of
19505	TICAM1	22124111	IFN-gamma. (Demonstrated in mice) TICAM1 forms a complex with RIPK3 upon Toll-like receptors (TLR) 3 and 4 activation resulting in RIPK3- dependent but TNF-independent necrosis in
19505	TICAM1	22123964	macrophages. (Demonstrated in mouse) TICAM1 (TRIF) enhances expression of Kaposi †s
19505	TICAM1	23723066	sarcoma-associated herpesviral protein RTA. The homotypic interaction of TICAM2 Toll/interleukin-1 receptor (TIR) domain is essential to form a scaffold for recruiting the TICAM1 TIR
19505	TICAM1	24255114	domain. MSR1 and Cd36 share several ligands and are involved in largely overlapping physiological and pathological processes, but they differ significantly in their effects on proinflammatory and
136838	Cd36	24257313	immunoregulatory functions of macrophages. MSR1 is required for sensing human cytomegalovirus (HCMV) by endosomal TLR3 and TLR9 in monocytic
8289	MSR1	19914718	THP-1 cells. MRS1 (SR-A) is upregulated in TLR4-mediated LPS responses and these receptors contribute to the efficient capturing and clearance of invading microbial
8289	MSR1	20162551	pathogens. MSR1 deficiency leads to greater sensitivity to LPS- induced endotoxic shock. MSR1 down-regulates inflammatory gene expression in dendritic cells by
8289	MSR1	21460221	suppressing TLR4-mediated activation of NFKB.

			MSR1, one of the principal receptors expressed on macrophages, suppresses macrophage activation by inhibiting the binding of lipopolysaccharide (LPS) to TLR4 in a competitive manner; thus playing a pivotal
8289	MSR1	21756882	role in the regulation of the LPS-induced inflammatory response. (Demonstrated in murine model) Deletion of MSR1 results in protection from septic
8289	MSR1	22751446	shock and modulation of TLR4 signalling in peritoneal macrophages. (Demonstrated in mice) MSR1 and Cd36 share several ligands and are involved in largely overlapping physiological and pathological processes, but they differ significantly in their effects on proinflammatory and
8289	MSR1	24257313	immunoregulatory functions of macrophages. HMGB1 functions as universal sentinel for nucleic-
21100	HMGB1	19890330	acid-mediated innate immune responses. HMGB1 activates innate immunity mechanisms as a
21100	HMGB1	19914413	<ul> <li>complex with DNA, lipids and/or pro-inflammatory cytokines.</li> <li>HMGB1 is an alarmin and a key mediator of natural killer (NK)-dendritic cell (DC) cross-talk and plays a pivotal role in the escape of Human Immunodeficiency Virus (HIV)-infected dendritic cells from TRAIL-mediated NK cell cytotoxicity during NK-DC cross-</li> </ul>
21100	HMGB1	20419158	talk by upregulating c-FLIP and c-IAP2 expression. HMGB1 is an endogenous TLR4 ligand in macrophages and its release in wounds initiates TLR4-
21100	HMGB1	21372296	dependent responses that contribute to neovascularization. (Demonstrated in murine model) HMGB1 is found in high concentrations within neutrophil extracellular traps (NETs). HMGB1 is a neutrophil protein that facilitates the uptake and recognition of mammalian DNA by plasmacytoid dendritic cells, and may play a role in Systemic Lupus
21100	HMGB1	21389264	Erthermatosus autoimmunity. HMGB1 plays a key regulatory role in polymorphonuclear neutrophil (PMN) recruitment to inflammatory tissues. Low concentrations of HMGB1 (50-100 ng/ml) reduce baseline PMN migration as well as formyl-methionyl-leucyl-phenylalanine- and IL8- induced PMN chemotaxis, whereas higher HMGB1
21100	HMGB1	21860212	concentrations (5000 ng/ml) have a chemoattractant effect on PMN through IL8 production. HMGB1 has a pathogenic role in arthritis, where in complex with lipopolysaccharide, IL1A or IL1B, HMGB1 boosts the production of proinflammatory cytokines and MMP3 as demonstrated in synovial fibroblasts from rhoumatoid arthritis and actaoarthritis
21100	HMGB1	21871094	fibroblasts from rheumatoid arthritis and osteoarthritis patients. Nuclear HMGB1 translocates to the cytoplasm in LPS- atimulated macrophages to potentiate inflammatory
21100	HMGB1	22396017	stimulated macrophages to potentiate inflammatory responses. (Demonstrated in mice)

			Mast cell chymase CMA1 contributes to the control of inflammation by degrading the virulence factor Hsp70 of Trichinella spiralis, as well as several alarmins such
21100	HMGB1	24257755	as endogenous HSPA1A, BGN, and HMGB1 Mast cell chymase CMA1 contributes to the control of inflammation by degrading the virulence factor Hsp70 of Trichinella spiralis, as well as several alarmins such
89505	BGN	24257755	as endogenous HSPA1A, BGN, and HMGB1 HSPA1A is secreted into the extracellular space during exercise-induced stress and increases the intracellular
79274	HSPA1A	21448922	levels of cAMP, which acts as an "intracellular danger signal" to activate neutrophils. Mast cell chymase CMA1 contributes to the control of inflammation by degrading the virulence factor Hsp70
79274	HSPA1A	24257755	of Trichinella spiralis, as well as several alarmins such as endogenous HSPA1A, BGN, and HMGB1. Mast cell chymase CMA1 contributes to the control of inflammation by degrading the virulence factor Hsp70 of Trichinella spiralis, as well as several alarmins such
3997	CMA1	24257755	as endogenous HSPA1A, BGN, and HMGB1 TIRAP is a Toll/Interleukin-1 receptor (TIR) domain containing adapter protein that binds to TLR4, serving
75811	TIRAP	11526399	as a bridge for MYD88 recruitment. TIRAP is dispensable in TLR2 signalling at high ligand concentrations in macrophages and dendritic cells, with MyD88 probably coupling to the TLR2 receptor complex at sufficient levels to allow activation
75811	TIRAP	19717524	but has an inhibitory role in the signalling of TLR3 to JNK.
75811	TIRAP	11544529	TIRAP is an adapter in Toll-like receptor 4 (TLR4) signal transduction. TIRAP is differentially involved in signalling by members of the Toll-like receptor (TLR) family and
75811	TIRAP	12447442	may account for specificity in the downstream signalling of individual TLRs.
75811	TIRAP	12447441	TIRAP has a crucial role in the MyD88-dependent signalling pathway shared by TLR2 and TLR4. TIRAP is a substrate for IRAK1 and IRAK4 with
75811	TIRAP	20400509	phosphorylation promoting its ubiquitination and degradation. TIRAP is an activator of TLR2/4 signalling and a negative regulator of TLR3/TRIF signalling. TIRAP is
75811	TIRAP	2095775	essential in restricting TLR3 signalling thereby protecting the host from unwanted immunopathologies associated with excessive IFN-beta production. TIRAP Ser180Leu polymorphism is significantly associated with Behcet's disease in UK, but not Middle Eastern, patients. It is suggested that the Ser180Leu functional variant of TIRAP will lead to greater
75811	TIRAP	21705416	cytokine production and tissue damage with persistence of mucosal lesions upon encounter with pathogens.

			TIR domain-contaning protein from Brucella melitensis, TcpB, disrupts the receptor-adaptor
75811	TIRAP	24265315	interaction between TLR4 and TIRAP. C3a and its receptor C3ar1 are critical for defense
193788	C3	24273177	against C. psittaci in mouse lung infection. C3a and its receptor C3ar1 are critical for defense
185906	C3ar1	24273177	against C. psittaci in mouse lung infection. Mycobacterial infection induces expression of Mir155, which promotes the maturation of phagosomes and
142234	Rheb	24130493	represses the expression of Rheb by targeting its 3†UTR The interplay between Tsc1-Rheb-mTOR complex 1
			signalling and Myc-dependent metabolism are dynamically regulated during dentritic cell (DC) development, whereas uncontrolled mTORC1
142234	Rheb	24282297	activation impairs DC development. Tsc1 inhibits TLR response and endotoxin tolerance
152579	Tsc1	22412198	through repression of mTORC1 and JNK1/2 signalling pathways.
152579	Tsc1	23776173	TSC1 is a critical regulator of dendritic cell function in both innate and adaptive immunity. The interplay between Tsc1-Rheb-mTOR complex 1 signalling and Myc-dependent metabolism are
152579	Tsc1	24282297	dynamically regulated during dentritic cell (DC) development, whereas uncontrolled mTORC1 activation impairs DC development. Inhibition of mTOR blocks the anti-inflammatory
204403	Mtor	21368289	potency of glucocorticoids both in human monocytes and myeloid dendritic cells. (Demonstrated in human) The interplay between Tsc1-Rheb-mTOR complex 1 signalling and Myc-dependent metabolism are
204403	Mtor	24282297	dynamically regulated during dentritic cell (DC) development, whereas uncontrolled mTORC1 activation impairs DC development. Mir126-Kdr axis is an important regulator of the innate
172240	Kdr	24270517	response. Mir126 controls the survival and function of plasmacytoid dendritic cells and regulates gene expression of Tlr7, Tlr9, Nfkb1 and Kdr. Mir126 controls the IFN- $\hat{I}\pm/\hat{I}^2$ responses to pathogen-
172240	Kdr	24270517	associated nucleic acids through regulating the homeostasis and function of plasmacytoid dendritic cells (pDCs). Mir126 operates in part by regulating Kdr which encodes the growth factor receptor VEGFR2 in pDCs Mir126-Kdr axis is an important regulator of the innate response. Mir126 controls the survival and function of
224249	Mir126	24270517	plasmacytoid dendritic cells and regulates gene expression of Tlr7, Tlr9, Nfkb1 and Kdr.

			Mir126 controls the IFN- $\hat{l}\pm/\hat{l}^2$ responses to pathogen- associated nucleic acids through regulating the homeostasis and function of plasmacytoid dendritic cells (pDCs). Mir126 operates in part by regulating Kdr which encodes the growth factor receptor VEGFR2 in
224249	Mir126	24270517	pDCs Stat4 is required for the generation of an effective
152917	Stat4	15356157	innate host defense against bacterial pathogens of the lung. Ity14 (Immunity to Typhimurium locus 14) mice with a mutation in Stat4 have an increased innate
152917	Stat4	24285835	susceptibility following sublethal invasive S. Typhimurium challenge. ARF6 has a pivotal role in TLR9-mediated immune
5864	ARF6	22170068	signaling by regulating the cellular uptake of CpG oligodeoxynucleotides. ARF6 regulates LPS internalization and LPS-induced
5864	ARF6	24297182	relocation of TICAM2 (TRAM), which is required for the MyD88-independent TLR4 signalling cascade. Dusp16 has a dual function in the innate immune system shaping the output of cytokines and other TLR- induced gene products by macrophages, most notably
194120	Dusp16	24311790	IL12, and regulating the responsiveness of bone marrow progenitor cells to the growth factor GM-CSF Central protein of the complement system; Classical, lectin and alternative pathways of complement all
21798	C3	16875735	converge at the activation of C3 yielding a diverse set of biological responses T cell-expressed CTSL1 cleaves C3 into active C3a
21798	C3	24315997	and C3b fragments mediating the intracellular and extracellular C3 activation in T cells.
74981	CTSL1	20926012	CTSL1 enhances angiogenesis by increasing extracellular matrix degradation and remodelling. T cell-expressed CTSL1 cleaves C3 into active C3a
74981	CTSL1	24315997	and C3b fragments mediating the intracellular and extracellular C3 activation in T cells. TRAF2 and TRAF5 are signal transducers for the TNF
92817	TRAF2	11479302	receptor superfamily and are involved in TNF-induced NF-kappaB activation and protection from cell death. TRAF2 plays a critical role in TNF signalling by directing the IKK complex to the membrane, promoting TRAF2 K63-linked ubiquitination, and
92817	TRAF2	19150425	positioning the IKKalpha and IKKbeta chains with the TAK1/TAB kinase. TRAF2 competes with TRAF6 for CD40 binding,
92817	TRAF2	20449947	thereby limiting the capacity of CD40 engagement to induce NF-kappaB activation in human lymphocytes.

			TRAF2 deficiency results in the accumulation of TNF- dependent, IL10-secreting neutrophils. Combined treatment of neutralizing antibodies against both TNF and IL10 substantially ameliorated the colitis
92817	TRAF2	22546736	phenotype in the Traf2 null mice. (Demonstrated in mice) GNB2L1 (RACK1) negatively regulates NF Î ° B
92817	TRAF2	24323043	activation by interacting with CHUK and IKBKB. The interaction interferes with the recruitment of the IKK complex to TRAF2. GNB2L1 (RACK1) negatively regulates NF $\hat{1}$ ° B activation by interacting with CHUK and IKBKB. The interaction interferes with the recruitment of the IKK complex to TRAF2.
63034	GNB2L1	24323043	complex to TRAF2.
199298	Rnasel	21190483	Rnasel cleaves RNA during viral infections and the cleavage products induces the RIG-I pathway and production of Ifnb gene. In addition, Rnasel is implicated in the protection of central nervous system against viral-induced demyelination. A broader role in innate immunity is suggested by involvement of Rnasel in cytokine induction and endosomal pathways that suppress bacterial infections.
199290	Kliaser	21170403	RNASEL contributes to innate immunity through
199298	Rnasel	24324683	regulating macrophage functions. Phosphorylation of the inflammasome adaptor Pycard (ASC) controls inflammasome activity through the formation of ASC specks. The NLRP3 and AIM2
148417	Mapk8	24185614	inflammasomes require Syk and Mapk8 (JNK) for their full activity . After viral infection, ELF4 binds to TMEM173
85951	ELF4	24185615	<ul><li>(STING) and induces type I interferon. ELF4 is critical for host antiviral defense.</li><li>Cytotoxic stress exemplified by ionizing radiation induces IFN-beta and enhances the expression of Dhx58 which in turn suppresses the interferon</li></ul>
211512	Dhx58	24434553	stimulated genes associated with cytotoxic stress by turning off the expression of IFN-beta. JAK1 physically associates with IL2RB and tyrosine phosphorylation of JAK1 is induced upon IL2
99443	JAK1	8041779	stimulation, suggesting that regulation of JAK1 may be linked to IL2 induced signal transduction. JAK1 and JAK2 are tyrosine kinases involved in the regulation of cell proliferation, differentiation, and survival and simultaneous activation of both JAK1 and JAK2 fusion proteins, but not either one alone, leads to the tyrosine phosphorylation of IL3RB, the activation of downstream signalling molecules, including STAT5, AKT, and MAPK, and the conferring of factor-
99443	JAK1	15988755	independent growth to IL-3-dependent Ba/F3 cells.

			TNE
			TNK1 is a component of the IFN-JAK-STAT signalling cascade and is a critical antiviral host factor where its abundance is inversely correlated to viral
99443	JAK1	24449862	replication and contributes to the hepatocytic response to antiviral treatment. TNK1 is a component of the IFN-JAK-STAT
			signalling cascade and is a critical antiviral host factor where its abundance is inversely correlated to viral
24740	TNK1	24449862	replication and contributes to the hepatocytic response to antiviral treatment.
			Reg3g has a protective role against mucosal infection by binding to Gram-negative and Gram-positive
158682	Reg3g	24345802	bacteria, and influences mucus distribution in the ileum and contributes to mucosal protection.
			Nlrx1 deletion leads to constitutive interaction of Mavs and Ddx58, resulting in increased inflammation. Nlrx1 knockout mice showed increased expression of
			antiviral signalling molecules such as Ifnb, Stat2, Oas1 and Il6 after influenza infection. In addition to attenuating the antiviral response, Nlrx1 directly
			associates with Traf6 and inhibits NFkB pathway in
156238	Nlrx1	21703540	LPS-activated macrophages. Nlrx1 negatively regulates TLR-mediated NFkB
			activation by directly interacting with Traf6 or IKK kinase. Nlrx1 knockdown in mice enhances their
156238	Nlrx1	21703539	susceptibility to LPS-induced septic shock and increases plasma Il6 levels.
150258	MIXI	21705559	Nlrx1 attenuates type I IFN production and promotes autophagy during viral infection. (Demonstrated in
156238	Nlrx1	22749352	human) Nlrx1 attenuates macrophage/microglial inflammatory
			activities and protects against experimental autoimmune encephalomyelitis caused by autoreactive
156238	Nlrx1	24366868	T cells.
			HIV auxiliary protein Vpr directly activates SLX4 to promote cell cycle arrest at the G2/M, a process that
12161	SLX4	24412650	requires VPRBP-DDB1-CUL4 E3-ligase complex, to escape from innate immune sensing.
			Microbiota-driven Il17c induces Bcl2 and Bcl211 expression in intestinal epithelial cells in an autocrine
			manner to promote cell survival and tumorigenesis in
209867	Bcl211	24412611	both chemically induced and spontaneous intestinal tumor models.
			Bcl2 is a multifunctional regulator of cell survival that inhibits the innate immune response during early stages of pathogenesis. Muscle-specific expression of Bcl2 in
			Lama2-deficient mice resulted in the inhibition of Tlr4, Tlr6, Tlr7, Tlr8 and Tlr9 induction, leading to reduced
185673	Bcl2	21850221	infiltration of eosinophils, the principal death effector cells.

			Microbiota-driven Il17c induces Bcl2 and Bcl2l1 expression in intestinal epithelial cells in an autocrine manner to promote cell survival and tumorigenesis in both chemically induced and spontaneous intestinal
185673	Bcl2	24412611	tumor models. Il17c is an essential autocrine cytokine that regulates innate epithelial immune responses. (Demonstrated in
197277	Il17c	21993848	human) Microbiota-driven Il17c induces Bcl2 and Bcl2l1 expression in intestinal epithelial cells in an autocrine manner to promote cell survival and tumorigenesis in both chemically induced and spontaneous intestinal
197277	Il17c	24412611	tumor models. Il27 priming results in an enhanced LPS-induced proinflammatory response in primary monocytes, specifically increasing expression of Il6, Tnf, Ccl3, and
209673	1127	22156348	Ccl4. (Demonstrated in human) Influenza virus infection induces pulmonary II27 production in a type I IFN-alpha/beta receptor (IFNAR) signalling-dependent manner, which sensitizes mice to secondary pneumococcal infection downstream of
209673	Il27	24408967	IFNAR pathway. Gata3 is a critical regulator of innate lymphoid cell
135340	Gata3	23733962	development from hematopoietic stem cells. Gata3 plays a generalized role in innate lymphoid cell (ILC) lineage determination and is critical for the
135340	Gata3	24419270	development of gut Rorc+ group 3 ILCs subsets that maintain mucosal barrier homeostasis. Regulation of IL1B-induced NF-ΰB by hydroxylases links key hypoxic and inflammatory signaling
728297	EGLN2	24145445	pathways in a manner that is dependent upon the combinatorial blockade of both EGLN2 and HIF1AN. Regulation of IL1B-induced NF-ΰB by hydroxylases links key hypoxic and inflammatory signaling nethodates in a manner that is dependent upon the
86613	HIF1AN	24145445	pathways in a manner that is dependent upon the combinatorial blockade of both EGLN2 and HIF1AN. Cyclic-di-GMP-induced levels of Ifi202b suppress the
206063	Ifi202b	24131791	expression of Tmem173 (STING). Il9 production is largely restricted to innate lymphoid cells during papain-induce lung inflammation and serves as an as important bridging link to induce type 2
157258	I19	21983833	helper T cell responses. Th9 cells and IL-9 have a critical and nonredundant
157258	119	24138883	role in host-protective type 2 immunity against parasitic worm infection A type-I-IFN receptor that functions as a nuclear
1833	CRKL	9872990	adapter protein and associates with STAT5 to regulate gene transcription through DNA binding Enterohemorrhagic E. coli NleH1 effector targets CRKL to inhibit ribosomal protein S3 nuclear
1833	CRKL	24145029	translocation and the NF <sup>î</sup> <sup>e</sup> B pathway.

			Nod2 driven inflammation is regulated by nitric oxide responsive Mir146 that facilitates activation of sonic hedgehog (SHH) signalling by targeting Numb
155803	Numb	24092752	expression. Vaccinia virus protein C16 influences the immune response by binding to the XRCC6/XRCC5 (Ku70/80)
80616	XRCC5	24098118	complex thus blocking PRKDC-dependent DNA sensing in fibroblasts Vaccinia virus protein C16 influences the immune response by binding to the XRCC6/XRCC5 (Ku70/80)
9323	XRCC6	24098118	complex thus blocking PRKDC-dependent DNA sensing in fibroblasts Vaccinia virus protein C16 influences the immune
408612	PRKDC	24098118	response by binding to the XRCC6/XRCC5 (Ku70/80) complex thus blocking PRKDC-dependent DNA sensing in fibroblast. Ifit1 recognizes and directly binds to 5'-triphosphate
159854	Ifit1	21642987	RNA and functions as an innate inhibitor of viral replication and pathogenicity. Ifit1 is an innate immune network bottleneck with the
159854	Ifit1	22745654	ability to suppress induction of Tlr4 response genes in LPS-stimulated macrophages. IFIT1 binds mRNAs that lack 2â€ <sup>2</sup> O methylation on
159854	Ifit1	24098121	the first ribose, thus inhibiting translation by impairing binding of eukaryotic translation initiation factors to 2â€ <sup>2</sup> O-unmethylated RNA templates Mfn2 deletion causes inability of the cell to undergo mitochondrial fusion, and therefore exhibit impaired induction of interferons and proinflammatory cytokines in response to viral infection - which results in
203813	Mfn2	21285412	increased viral replication.
203813	Mfn2	24127597	Mitochondrial membrane potential is required for the association of Nlrp3 and Mfn2. Mfns2 is required for the activation of Nlrp3 inflammasomes. F2RL1 and TLR4 interact in a synthetic agonist peptide-dependent manner and may support a novel
29713	F2RL1	18622013	paradigm of receptor cooperativity in inflammatory responses.
29713	F2RL1	19675284	<ul><li>F2RL1 is involved in inflammation and is specifically activated by mast cell-derived tryptase.</li><li>Activation of F2RL1 stimulates IL6, IL8, and PTGER2 (prostaglandin E2) release from human respiratory</li></ul>
29713	F2RL1	11907122	epithelial cells, thus having an important role in modulating inflammation in the lung. F2RL1 cooperates with TLR2, TLR3, or TLR4 for
29713	F2RL1	19865078	activation of nuclear factor-kappaB-dependent signalling in mucosal epithelial cell lines. Activation of F2RL1 negatively regulates TLR3- dependent antiviral pathway, blunting the expression of
29713	F2RL1	19865078	TLR3/IRF3 driven genes, as well as activation of IRF3 and STAT1.
			F2RL1 (PAR2) activates MAPK14 (p38) to trigger production of innate immunity markers in oral
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29713	F2RL1	21029417	kertainocytes. IL32 interacts with PR3 and F2RL1, triggering the PAR2-TRIF signalling axis, and IL32 may have a
29713	F2RL1	24129891	potential role in the transition from innate to adaptive immunity. IL32 interacts with PRTN2 (PR3) and F2RL1, triggering the PAR2-TRIF signalling axis, and IL32
13084	PRTN3	24129891	may have a potential role in the transition from innate to adaptive immunity. Msr1 deficiency leads to greater sensitivity to LPS-
149582	Msr1	21460221	induced endotoxic shock. Msr1 down-regulates inflammatory gene expression in dendritic cells by suppressing Tlr4-mediated activation of NFKB. Msr1, one of the principal receptors expressed on
			macrophages, suppresses macrophage activation by inhibiting the binding of lipopolysaccharide (LPS) to Tlr4 in a competitive manner; thus playing a pivotal
149582	Msr1	21756882	role in the regulation of the LPS-induced inflammatory response.
149582	Msr1	24035364	Autophagy regulates phagocytosis by modulating the expression of scavenger receptors, Marco and Msr1
188086	Marco	24035364	Autophagy regulates phagocytosis by modulating the expression of scavenger receptors, Marco and Msr1 CEBPB, one of the CEBP family members, is a crucial
			regulator of gene expression during innate immunity, inflammatory responses and adipogenesis. All of the individual CEBPB isoforms, LAP1, LAP2 and LIP,
80858	CEBPB	18820298	attenuate EGF-induced PTGS2 (COX-2) promoter activity.
			CEBPB is a member of the CCAAT enhancer binding protein family and is a transcriptional factor regulating genes in innate immunity and inflammation. The
80858	CEBPB	21326902	activities of CEBP are regulated via methylation of arginine and lysine side chains.
80838	CEDFD	21320902	CREB1 and CEBPB are involved in cytokine
80858	CEBPB	24038085	production in neutrophils in response to LPS and TNF. CREB1 and AP1 transcription factor activation regulates LPS- or M-CSF-induced activation of the MAPK phosphatase-1 (MKP-1) gene, a protein
79573	CREB1	19585511	<ul><li>phosphatase that plays a crucial role in innate immunity.</li><li>CREB1 has a key role in transcriptional regulation of macrophage migration inhibitory factor (MIF) gene</li></ul>
79573	CREB1	18034423	expression and MIF-dependent host antimicrobial innate immune defence.
79573	CREB1	24038085	CREB1 and CEBPB are involved in cytokine production in neutrophils in response to LPS and TNF.

Nos2 (iNOS) and Calm1 (CaM) coordinately function to form a stable complex that is part of a rapid host response that functions within the first 30 min following bacterial infection to upregulate the innate immune system involving macrophage activation.

203203	Nos2	16893173	immune system involving macrophage activation.
203203	Nos2	22542147	Nitric oxide production by Nos2 promotes Listeria monocytogenes dissemination in the host. Inorganic polyphosphate suppresses Nos2 expression
203203	Nos2	24040305	and nitric oxide release but not TNF release in LPS- activated macrophages IL28A is able to efficiently inhibit Herpes Simplex Virus Type-1 replication in neuronal cells by inducing
50102	IL28A	21499846	the expression of TLR as well as activate the TLR- mediated interferon antiviral pathway. IFNL1, IFNL2 (IL28A) and IFNL3 have different effects on Toll-like receptor-related gene expression in
50102	IL28A	24041672	HepG2 cells. Nfe2l2 plays an important role in Tlr4-mediated autophagy. Nfe2l2 is activated by reactive oxygen species-Mapk14 axis-dependent Tlr4 signaling, and induces the accumulation of Sqstm1 and aggresome- like induced structures.
181483	Nfe2l2	21220332	
181483	Nfe2l2	24011591	Phosphorylation of Sqstm1 (p62) activates the Keap1- Nrf2 pathway during selective autophagy Phosphorylation of Sqstm1 (p62) activates the Keap1-
139531	Keap1	24011591	Nrf2 pathway during selective autophagy IL12B production in response to Toll-like receptor (TLR) stimulation is regulated by the tyrosine phosphatase activity of PTPN6 and this is a novel mechanism for host regulation of IL12, a cytokine
55893	IL12B	20145200	important in both innate and adaptive immunity. IL12, consisting of IL12A and IL12B subunits, initiates local antitumor immunity by stimulating lymphoid tissue-inducer (LTi) cells bearing the natural
55893	IL12B	20935648	cytotoxicity receptor NCR1 (NKp46). IL12, consisting of IL12A and IL12B subunits, induces IL2RA to form high-affinity IL2 receptors on natural killer cells in response to mouse cytomegalovirus
55893	IL12B	22888135	infection. (Demonstrated in mouse) Macrophages cultured with bile acids produce lower levels of IL12 and display an anti-inflammatory phenotype characterized by an increased ratio of IL10
55893	IL12B	23990628	to IL12 IFN gamma creates a primed chromatin environment in
55893	IL12B	24012417	macrophages to augment TLR-induced IL12B transcription Cebpb is a member of the CCAAT enhancer binding protein family and is a transcriptional factor regulating genes in innate immunity and inflammation. The activities of CEBP are regulated via methylation of
213090	Cebpb	21326902	arginine and lysine side chains.

			LPS stimulation promotes the formation of CEBPB complexes on the Serpinb2 promoter to drive
213090	Cebpb	23472114	transcription. The serpino2 promoter to drive
213090	Cebpb	24023826	Cebpb in response to LPS in macrophages. TRIM25 is essential for RIG-I-mediated antiviral activity by inducing Lys 63-linked ubiquitination of
60223	TRIM25	17392790	DDX58 which is crucial for RIG-I signalling pathway. TRIM25 was identified in a systematic screen for
60223	TRIM25	23438823	positive regulators of innate immune responses. RNF135 is essential for the association of DDX58 (RIG-I) and TRIM25, resulting in the activation of
60223	TRIM25	23950712	RIG-I signalling. RNF135 is essential for the association of DDX58 (RIG-I) and TRIM25, resulting in the activation of
39370	RNF135	23950712	RIG-I signalling. IKBKG is the regulatory subunit of the IKK- signalosome complex that is required for
777831	IKBKG	10968790	proinflammatory activation of the I-kappaB-kinase (IKK) complex. IKBKG is linearly polyubiquitinated by the LUBAC ligase complex, consisting of RNF31 and RBCK1
			proteins, to regulate the canonical NF-kappaB activation pathway.
777831	IKBKG	19136968	IKBKG regulates TNF alpha signaling by coordinating
777831	IKBKG	18080803	cell responses mediated by the AP-1 and NF-kappa B pathways. IKBKG is essential for NF-kappaB activation and the polyubiquitylation and the degradation of IKBKG
777831	IKBKG	20010814	during Shigella infection is a new bacterial strategy to modulate host inflammatory responses. The binding of MAVS to Traf2, Traf5, and Traf6 is
777831	IKBKG	23951545	dependent on virus infection and MAVS polymerization. The TRAF proteins promote ubiquitination that recruits IKBKG binding to the MAVS signalling complex The binding of MAVS to Traf2, Traf5, and Traf6 is dependent on virus infection and MAVS
208924	Traf5	23951545	polymerization. The TRAF proteins promote ubiquitination that recruits IKBKG binding to the MAVS signalling complex
205458	Mertk	23954153	Viruses engage Tyro3/Axl/Mertk (TAM) receptors in order to inhibit host type I IFN signalling.
158468	Axl	23954153	Viruses engage Tyro3/Axl/Mertk (TAM) receptors in order to inhibit host type I IFN signalling.
199993	Tyro3	23954153	Viruses engage Tyro3/Axl/Mertk (TAM) receptors in order to inhibit host type I IFN signalling. LGALS9 attenuates acute lung injury by preferentially suppressing pro-inflammatory functions in pDC-like
35872	LGALS9	21562126	macrophages through TLR2/TLR4 down-regulation.

35872	LGALS9	23408620	LGALS9 engagement impairs the cytotoxicity function and cytokine production of natural killer cells. TLR activation promotes HAVCR2 and LGALS9 association within the same macrophage to
35872	LGALS9	23967307	<ul><li>differentially regulate IL12/IL23 expressions via STAT3 phosphorylation.</li><li>CLEC4D and CLEC6A form a heterodimer complex which confers innate cells high ability to sense C.</li></ul>
17159	CLEC4D	23911656	albicans infection by facilitating the activation of NFÎ <sup>o</sup> B dependent inflammatory responses. MIR145 directly targets HDAC11 to promote IL10
19657	HDAC11	23980205	expression in TLR4-triggered macrophages. MIR145 targets SOCS7 to promote IFNB induction in
126739	MIR145	23392170	bladder cancer cells. MIR145 directly targets HDAC11 to promote IL10
126739	MIR145	23980205	expression in TLR4-triggered macrophages PARK2 was recently shown to promote the clearance of impaired mitochondria by autophagy, termed mitophagy; PARK2 promotes mitophagy by catalyzing mitochondrial ubiquitination, which in turn recruits ubiquitin-binding autophagy components, HDAC6 and
98830	PARK2	20457763	p62, leading to mitochondrial clearance. PARK2 is important for innate defence against M. tuberculosis for having a role in ubiquitin-mediated
98830	PARK2	24005326	autophagy of M. tuberculosis During the transcriptional response to Sendai virus
7203	POLR2F	23994473	infection, POLR2F(RNA Pol II) is recruited by IRF3 and NF PB to control virus induced gene activation. IDO1 limits innate and adaptive immunity to apoptotic self-antigens. IDO1-mediated inhibition of inflammation plays a key role in suppressing systemic
18750	IDO1	22355111	autoimmune diseases. (Demonstrated in mice) DNA nanoparticle is sensed selectively by myeloid dendritic cells (DCs) via the STING (for stimulator of interferon genes) /type I IFN pathway to induce Ido1 in
18750	IDO1	23986532	DCs, which activate regulatory T cells. MIR141 directly regulates CXCL12 expression and
71595	CXCL12	24000293	CXCL12 mediated leukocyte migration during colonic inflammation. MIR141 directly regulates CXCL12 expression and
126529	MIR141	24000293	CXCL12 mediated leukocyte migration during colonic inflammation MIR21 inhibition enhances CCL5 (RANTES) and CXCL10 (IP-10) release in MCF-7 cancer cells and
101753	PIAS3	23998932	resulted in increased lymphocyte migration. PIAS3 is a target of MIR21 in MCF-7 cells. CCL5 expression activates the IFN response during C. pneumoniae infection is mediated by intracellular nucleotide-sensing pattern recognition receptors (PRRs), which operate through a mechanism dependent
42204	CCL5	20386592	on the bacterial type III secretion system (T3SS).

			CCL5 expression in airway epithelial cells is regulated by cross-talk between TLR3 signalling and
42204	CCL5	20523058	inflammatory cytokines where TNF-alpha activates NF-kappaB, in cooperation with TLR3 signalling. MIR21 inhibition enhances CCL5 (RANTES) and CXCL10 (IP-10) release in MCF-7 cancer cells and
42204	CCL5	23998932	resulted in increased lymphocyte migration. PIAS3 is a target of MIR21 in MCF-7 cells. MIR21 targets PDCD4, which acts a molecular switch between the pro-inflammatory (NFkB) and anti- inflammatory (IL10) response. MIR21-mediated reduction of PDCD4 is essential for protecting against
126673	MIR21	21652514	the lethal effects of LPS. MIR21 is upregulated during Mycobacterium leprae infection of monocytes to escape from vitamin D-
126673	MIR21	22286305	dependent antimicrobial pathways.
126673	MIR21	23580661	Circulating MIR21 activates natural killer cells via the TLR1 signalling pathway. MIR21 inhibition enhances CCL5 (RANTES) and
			CXCL10 (IP-10) release in MCF-7 cancer cells and resulted in increased lymphocyte migration. PIAS3 is
126673	MIR21	23998932	a target of MIR21 in MCF-7 cells. MIR133A1 enhances CASP1 activation and IL1B
64056	UCP2	23988448	processing and suppresses inflammasome activation by suppressing its target UCP2 MIR133A1 enhances CASP1 activation and IL1B
127431	MIR133A1	23988448	processing and suppresses inflammasome activation by suppressing its target UCP2 IL12, consisting of IL12A and IL12B subunits,
63368	IL12A	20935648	initiates local anti-tumour immunity by stimulating lymphoid tissue-inducer (LTi) cells bearing the natural cytotoxicity receptor NCR1 (NKp46). IL12, consisting of IL12A and IL12B subunits, induces IL2RA to form high-affinity IL2 receptors on natural
63368	IL12A	22888135	killer cells in response to mouse cytomegalovirus infection. (Demonstrated in mouse) Macrophages cultured with bile acids produce lower
63368	IL12A	23990628	levels of IL12 and display an anti-inflammatory phenotype characterized by an increased ratio of IL10 to IL12 RNF125 is an ubiquitin ligase which negatively
2177	RNF125	17460044	regulates RIG-I signalling by conjugating ubiquitin to DDX58 and MDA5 and IPS1, tagging them for proteosomal degradation
2177	RNF125	23772026	RNF125 is inhibited by human Bocavirus protein, VP2, to induce the IFNB signalling pathway. ZBTB20 inhibits transcription of IkBa to promote full
161026	Zbtb20	23776228	TLR-triggered immunity. Zbtb20 null mice are resistant to endotoxin shock and sepsis.
94666	ASCC3	23781071	ASCC3 inhibits IFN-signalling to dampen antiviral innate immunity.

			MASP2 binds to and is activated by MBL or ficolin in
89176	MASP2	18596036	response to pathogen-associated molecular patterns (PAMPs). MASP2, upon activation, can cleave complement
			component 4 (C4) and C2 to generate the C3
89176	MASP2	16189649	convertase, C4bC2b. MASP2 deficient mice are defective in the lectin
			pathway of complement activation and are highly
			susceptible to pneumococcal infection. (Demonstrated
89176	MASP2	22792067	in mice)
89176	MASP2	23785123	MASP2 forms a complex with MASP1 to activate the complement system.
0,1,0		20,00120	MASP1 is an essential protease of both the lectin and
			alternative complement pathways, essential
			components of innate immunity, participating in the pathogenesis of inflammatory diseases and in host
69257	MASP1	20038603	defence.
			MASP1 is crucial for classical complement activation,
69257	MASP1	22966085	but is not required for the alternative pathway function.
69257	MASP1	23785123	MASP1 forms a complex with MASP2 to activate the complement system.
0,231		25705125	Secretion of WNT9B upon virus infection negatively
55716	WNT9B	23785285	regulates antiviral innate immunity.
101110	WAITOD	22785285	Secretion of WNT2B upon virus infection negatively
101110	WNT2B	23785285	regulates antiviral innate immunity. Mutations in VPS45 are associated with congenital
102045	VPS45	23738510	neutrophil defect syndrome.
			Upon infection with encephalitic Bunyavirus, SARM1
202876	Sarm1	23499490	is activated by RIG-I/MAVS signalling to mediate neuronal cell death.
202870	Saliii	23499490	Sarm1 is expressed in neurons and regulates the
202876	Sarm1	23751821	expression of inflammatory and anti-viral cytokines.
	~ .		Sarm1 null mice are resistant to central nervous system
202876	Sarm1	23749635	infection with vesicular stomatitis virus. TLR9 acts as a receptor for unmethylated CpG-DNA
			and bacterial DNA to initiate a variety of immune
407857	TLR9	11130078	responses.
			TLR9 recognizes unmethylated DNA with CpG-motifs
			and the LR9 mediated signalling pathway is not only responsible for activation of innate immune cells, but
407857	TLR9	18262306	also for mounting acquired responses.
			TLR9, 8 and 7 form a functional subgroup within the
			TLR family that recognizes pathogen-associated molecular patterns in endosomal/lysosomal
407857	TLR9	14579267	compartments.
			TLR9 acts at the cell surface and engages an
107057	τι do	11067600	intracellular signalling pathway that includes MyD88,
407857	TLR9	11867692	IRAK, and TRAF6.

TLR9/7-me	ediated	innate	immune	responses	are
negatively	regulate	d via se	lected TLR	pathways	by a
human	mic	rosatelli	te I	DNA-mimi	cking
oligodeoxy	nucleoti	de with <b>(</b>	CCT repeat	S.	

TLR9 and TLR4 have both non-redundant and cooperative roles in lung innate responses during Gram-negative bacterial pneumonia and are both critical for IL-17 driven antibacterial host response.

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TLR9

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TLR9 ectodomain is cleaved to generate a functional receptor; although both the full-length and cleaved forms of TLR9 are capable of binding ligand, only the processed form recruits MyD88 on activation, indicating that this truncated receptor, rather than the full-length form, is functional.

TLR9 signalling enhances the rate of acidification of Salmonella-containing phagosomes, and this acidification induces the expression of Salmonella pathogenecity genes that are necessary for intracellular survival, growth, and systemic infection. TLR9 deficiency rescues the high Salmonella susceptibility phenotype observed in TLR2,TLR4 double mutant mice . (Demonstrated in murine model)

TLR9 triggers plasmacytoid dendritic cells in Systemic Lupus Erythematosus patients upon recognition of selfantigens such as neutrophil extracellular traps (NETs).

TLR9 requires proteolytic processing in endolysosome by asparagine endopeptidase and cathepsin in the endolysosome to initiate signalling in response to 21402738 DNA. (Demonstrated in murine model)

TLR9 deficiency reduced pancreatic edema, inflammation and pro-IL1B expression in pacreatitis. (Demonstrated in murine model)

TLR9 synergistically interacts with TLR2::TLR6 in lung epithelium to induce rapid pathogen killing, and can be used as a therapeutic target to treat otherwise lethal pneumonia.

> TLR9 is proteolytically cleaved in the endosome to form a soluble TLR9 (sTLR9), which inhibits TLR9dependent signalling and contributes to the prevention of autoimmune disease.

TLR9 activation is enhanced by increased levels of circulating histones, serving as a crucial link between initial damage and activation of innate immunity during sterile inflammation. (Demonstrated in murine model)

TLR9 promotes macrophage HIF1A levels, oxidative<br/>burst and nitric oxide production in response to group<br/>A Streptococcus (GAS), contributing to GAS clearance<br/>in vivo in both localized cutaneous and systemicTLR921860217TLR921860217

			TLR9 is selectively compartmentalized to fungal
407857	TLR9	21947771	phagosomes and negatively modulates macrophage anti-fungal effector functions. (Demonstrated in mice) TLR9 expression and signalling capacity oscillate with
407857	TLR9	22342842	the circadian clock. (Demonstrated in mice) Mitochondrial DNA that escapes from autophagy induces TLR9 inflammatory responses in
407857	TLR9	22535248	cardiomyocytes and is capable of inducing myocarditis and dilated cardiomyopathy. (Demonstrated in mice) TLR9 activation by endogenous self-ligands generated during oxidative stress promotes platelet hyper-
407857	TLR9	23071157	reactivity and thrombosis. Human papillomavirus E7 protein forms a transcriptionally repressive complex over the promoter
407857	TLR9	23752229	of TLR9 to suppress the interferon response. ANKRD17 binds to NOD2 and contributes to
23294	ANKRD17	23711367	inflammatory responses against Shigella flexneri. Human papillomavirus exploits UCHL1 to suppress interferon, pro-inflammatory cytokines and chemokines
15224	UCHL1	23717208	production and to dampen NF-kB signalling. KLF4 mediates the repression of Ciita/MHC-II in M. bovis BCG infected macrophages to evade antigen
148689	Klf4	23733190	presentation. ERN1 (IRE1)/XBP1-mediated signalling plays roles in the coordination of metabolic and immune responses by acting as a regulatory hub, linking endoplasmic
64539	ERN1	20533428	reticulum homeostasis with innate immunity and metabolism. Hyperactivated ERN1 (IRE1 $\hat{I} \pm$ ) increases TXNIP mRNA stability by reducing levels of a TXNIP destabilizing microRNA, miR-17. In turn, elevated TXNIP protein activates the NLRP3 inflammasome,
64539	ERN1	22883233	causing procaspase-1 cleavage and interleukin $1\hat{I}^2$ (IL- $1\hat{I}^2$ ) secretion. ERN1 binds to cholera toxin in the endoplasmic
64539	ERN1	23684307	reticulum to activate RIG-I innate immune signalling. COCH is secreted by follicular dendritic cells to
138038	Coch	23684986	promote antibacterial innate immunity. Cdc42 has a critical role in mediating innate immunity
198471	Cdc42	23690402	against upper airway infections. SRC and STAT3 play a role in apoptotic cell-mediated MerTK-dependent immunoregulation of dendritic cells.
74081	SRC	19667404	CD14 co-receptor participates with toll-like receptors (TLRs) in the response of microglial cells to fibrillar forms of bota amulaid contributing to microglial
49157	CD14	19776284	forms of beta-amyloid, contributing to microglial activation.

			CD14 recognizes necrotic cells in addition to LPS, PG, apoptotic cells, and lipids, suggesting that it might be a universal adaptor for damage-associated molecular
49157	CD14	19840871	pattern (DAMP) and pathogen-associated molecular patterns (PAMP). CD14 is a molecule that binds to lipopolysaccharide
49157	CD14	10652232	(LPS) and facilitates its signalling by helping TLR4- LY96 to sense and signal the presence of LPS. CD14 contributes to nucleic acid uptake in macrophages and acts as a co-receptor for endosomal TLR7/TLR9 activation.
49157	CD14	21078886	
49157	CD14	23548899	CD14 acts as an adaptor molecule for the immune recognition of salmonella curli fibers. IL27, a member of the IL-12 cytokine family, acts as a pro-inflammatory cytokine that regulates the
22826	IL27	20435892	differentiation of naive T helper cells and also possesses anti-inflammatory properties. IL27 induces a STAT1/3 and NFkappaB dependent pro-inflammatory cytokine profile in human
22826	IL27	20519510	monocytes. IL27 is a strong inducer of pro-inflammatory cytokine and chemokine expression, including enhancement of IL6, CXCL10 (IP-10), CCL3 (MIP-1alpha), CCL4 (MIP-1beta), and TNF (TNF-alpha) expression in human primary monocytes, the production of which is mediated by STAT1, STAT3, and NF-kappaB
22826	IL27	20519510	activation. IL27 priming results in enhanced LPS-induced proinflammatory response in human primary monocytes, specifically increasing expression of IL6,
22826	IL27	22156348	TNF, CCL3, and CCL4. IL27 reduces vacuolar ATPase levels to inhibit
22826	IL27	23557795	phagosome acidification in macrophages. Circulating MIR15B activates natural killer cells via
127561	MIR15B	23580661	the TLR1 signalling pathway. Lgr4 deletion potentiates TLR2/4-mediated cytokine
195566	Lgr4	23589304	<ul><li>production and Lgr4 null mice are more susceptible to septic shock.</li><li>TNFSF10 is targeted by human cytomegalovirus</li></ul>
65161	TNFSF10	23498957	glycoprotein to protect infected cells from natural killer cell cytotoxicity. PPP1CC dephosphorylates RNA sensors, RIG-I
197968	Ppp1cc	23499489	(DDX58) and MDA5 (IFIH1), to induce anti-viral IFNB production. PPP1CC dephosphorylates RNA sensors, RIG-I
57303	PPP1CC	23499489	(DDX58) and MDA5 (IFIH1), to induce anti-viral IFNB production. PPP1CA dephosphorylates RNA sensors, RIG-I
129195	Ppp1ca	23499489	(DDX58) and MDA5 (IFIH1), to induce anti-viral IFNB production.

60210	PPP1CA	18949366	PPP1CA positively regulates the TNF-alpha-induced NF-kappaB pathway at the level of IKK activation. PPP1CA dephosphorylates RNA sensors, RIG-I
60210	PPP1CA	23499489	(DDX58) and MDA5 (IFIH1), to induce anti-viral IFNB production. Mir203 post-transcriptionally downregulates Myd88 to
221154	Mir203	23522925	inhibit the TLR signalling pathway in macrophages. PSMB8 (LMP7) inhibits phagocytosis and enhances
170702	Psmb8	23527234	susceptibility of red blood cells to malarial parasites. MIR517C activates NF-kB signalling and directly
127401	MIR517C	23448136	targets TNIP1 mRNA to promote apoptosis. MIR517A activates NF-kB signalling and directly
127271	MIR517A	23448136	targets TNIP1mRNA to promote apoptosis. IFNE is constitutively expressed by epithelial cells of
162815	Ifne	23449591	the female reproductive tract to confer protection against sexually transmitted infections. CSF2 is a negative regulator of IL21-mediated
172875	Csf2	23453633	apoptosis in conventional dendritic cells.
140267	II21	23453633	IL21 induces apoptosis of conventional dendritic cells. IKBKE interacts with NKFB2 (p52) and promotes transactivation via RELA (p65)-regulation of the
776557	IKBKE	17003035	alternative NF-kappaB activation pathway involving p52 and p65. IKBKE is an IkappaB kinase that is expressed mainly in immune cells, and is induced in response to pro-
776557	IKBKE	10421793	inflammatory cytokines such as tumour necrosis factor (TNF)-alpha, IL-1-beta and IL-6, in addition to LPS. IKBKE and TBK1 have a pivotal role in coordinating
776557	IKBKE	12692549	<ul><li>the activation of IRF3 and NF-kappaB in the innate immune response.</li><li>IKBKE and other IKK kinases regulate each other by an intricate network involving phosphorylation of their</li></ul>
776557	IKBKE	21138416	catalytic and regulatory (NEMO, TANK) subunits to balance their activities during innate immunity.
			IKBKE kinase modulates IL-17 signalling through TRAF3IP2 adaptor protein phosphorylation, resulting in the induction of neutrophilia and pulmonary
776557	IKBKE	21822257	inflammation. (Demonstrated in mouse) IKBKE is an important signalling molecule in the IFN
776557	IKBKE	22065572	pathway and is necessary to mount anti-viral immunity against West Nile virus. (Demonstrated in mice) IKBKE plays a critical role in regulating the balance
776557	IKBKE	22171011	between the type I and type II interferon (IFN) signalling pathways. IKBKE is sequestered by arenavirus nucleoproteins to
776557	IKBKE	22532683	block its autocatalytic activity and its ability to active IRF3. IKBKE is poly-ubiquitinated at Lys30 and 401 to
776557	IKBKE	23453969	promote NF-kB activation in macrophages stimulated with TNFA and IL1B.

193646	Mfge8	23454767	MFGE8 inhibits inflammasome-induced IL1B production in macrophages. NOD1 and NOD2 represent central players in the
11085	NOD1	17690884	control of the immune responses to bacterial infections and inflammation. NOD1 and NOD2 can induce CCL5 (RANTES) through NF-kappaB pathway, orchestrating the global
11085	NOD1	17705131	Nod-dependent immune defence during bacterial infections. NOD1 and NOD2 direct autophagy by recruiting
11085	NOD1	19898471	ATG16L1 to the plasma membrane at the site of bacterial entry. NOD1 plays an important role in host defence against
11085	NOD1	20039881	bacterial infection by regulating direct killing of Helicobacter pylori bacteria by antimicrobial peptides. NOD1-dependent responses account for host resistance
11085	NOD1	20042586	against T. cruzi infection by mechanisms independent of cytokine production. NOD1 plays a role in priming innate defences,
11085	NOD1	20081863	facilitating a rapid response to infection by recognizing peptidoglycan from microbiota and enhancing killing of pathogens by bone marrow-derived neutrophils. NOD1 is a peripheral peptidoglycan intracellular sensor and is important for the progression and pathogenesis
11085	NOD1	21236705	of experimental autoimmune encephalomyelitis (animal model of multiple sclerosis).
			NOD1 detects heat-killed Legionella pneumophila and stimulates NFkB and IFN-beta promoter activity. NOD1 deficiency results in impaired bacterial clearance and increased proinflammatory cytokine at
11085	NOD1	21108472	24hrs post-infection. (Demonstrated in murine model) NOD1 is expressed by trophoblast cells across gestation and may have a role in mediating infection- associated inflammation and prematurity. Activation of NOD1 by bacterial peptidoglycan-derived peptide
11085	NOD1	21677137	induces maternal-fetal inflammation and preterm labour. Nod1 KO mice were protected from high-fat diet induced inflammation, lipid accumulation, and peripheral insulin intolerance. Ex vivo, NOD1 activation by bacterial peptidoglycan mimetics induces proinflammatory cytokine secretion and impaired insulin-stimulated glucose uptake in adipocytes. Hence, NOD1 is a plausible, new link between innate immunity and metabolism. (Demonstrated in murine
11085	NOD1	21715553	model)
11085	NOD1	23460743	Helicobacter pylori infection of gastric epithelial cells activates NOD1 to enhance IFN-gamma signalling. MIR3148 modulates allelic expression of TLR7
705497	MIR3148	23468661	variant, rs3853839G, associate with systemic lupus erythematosus.

			Plunc is a protein able to inhibit Mycoplasma pneumoniae (Mp) growth and its production following
210271	Divers	21054962	Mp infection is regulated through Toll-like receptor 2 (TL D2) signalling
210271	Plunc	21054862	(TLR2) signalling. Plunc is a secretory protein that exhibits antimicrobial
			activity against Gram-negative bacteria and anti-
210271	Plunc	21787346	inflammatory functions in respiratory infections. (Demonstrated in human)
210271	1 fulle	21/0/5/0	PLUNC1 is suppressed in pneumonic epithelium in the
210271	Plunc	23470624	respiratory tract. LPS stimulation promotes the formation of CEBPB
			complexes on the Serpinb2 promoter to drive
186363	Serpinb2	23472114	transcription.
			SFTPA1B;SFTPA1 is a carbohydrate pattern recognition molecule of innate immunity that
			recognition molecule of innate immunity that significantly enhances phagocytosis and killing of
			Aspergillus fumigatus, a pathogenic fungus, by
			neutrophils and macrophages.
80144	FTPA1B;SFTP	20413160	
			SFTPA1B;SFTPA1 can attenuate bacterial and viral
80144	FTPA1B;SFTP2	20418258	infection and inflammation by acting as an opsonin and by regulating innate immune cell functions.
00144	TIAID, SI TI I	20418238	SFTPA1 binds to Mycobacterium avium lipid and it
			promotes the agglutination of the pathogen. The
			presence of SFTPA1 inhibits the growth of M. avium
80144	FTPA1B;SFTP	21821801	in culture.
			SPA4 peptide, derived from SFTPA1, associates with
80144	FTPA1B;SFTP2	23475791	TLR4 to inhibit LPS-induced inflammation and to alleviate endotoxic shock.
00144	TIAID, SI TI I	23473791	TRIM63 was identified in a systematic screen for
94466	TRIM63	23438823	positive regulators of innate immune responses.
			TRIM61 was identified in a systematic screen for
43425	TRIM61	23438823	positive regulators of innate immune responses.
12 170		00.400.000	TRIM60 was identified in a systematic screen for
43470	TRIM60	23438823	positive regulators of innate immune responses.
23653	TRIM55	23438823	TRIM55 was identified in a systematic screen for positive regulators of innate immune responses.
23033	11(11)100	23 130025	TRIM49 was identified in a systematic screen for
67515	TRIM49	23438823	positive regulators of innate immune responses.
			TRIM45 was identified in a systematic screen for
101433	TRIM45	23438823	positive regulators of innate immune responses.
58977	TRIM42	23438823	TRIM42 was identified in a systematic screen for positive regulators of innate immune responses.
309//	1 K119142	23430023	TRIM38 targets TICAM1 (TRIF) for degradation to
66673	TRIM38	23056470	negatively regulateTLR3-mediated IFNB signalling.
			TRIM38 was identified in a systematic screen for
66673	TRIM38	23438823	positive regulators of innate immune responses.
(1551		22420022	TRIM37 was identified in a systematic screen for
61551	TRIM37	23438823	positive regulators of innate immune responses. TRIM36 was identified in a systematic screen for
37599	TRIM36	23438823	positive regulators of innate immune responses.

			TRIM27 was identified in a systematic screen for
72253	TRIM27	23438823	positive regulators of innate immune responses.
43164	TRIM24	23438823	TRIM24 was identified in a systematic screen for positive regulators of innate immune responses.
43104	I KIWI24	23430023	TRIM23 mediates K(lys)-27-ubiquitin conjugation to
			IKBKG (NEMO) which is essential for TLR3- and
			RIG-I/MDA5-mediated antiviral innate and
24045	TRIM23	20724660	inflammatory responses.
• • • • •			TRIM23 was identified in a systematic screen for
24045	TRIM23	23438823	positive regulators of innate immune responses.
27325	TRIM6	23438823	TRIM6 was identified in a systematic screen for positive regulators of innate immune responses.
21323	IKIMU	23430023	MID2 was identified in a systematic screen for positive
81697	MID2	23438823	regulators of innate immune responses.
			TRIM67 was identified in a systematic screen for
107395	TRIM67	23438823	positive regulators of innate immune responses.
			TRIM66 was identified in a systematic screen for
30708	TRIM66	23438823	positive regulators of innate immune responses.
(0001		22420022	TRIM65 was identified in a systematic screen for
69081	TRIM65	23438823	positive regulators of innate immune responses. TRIM56 is a regulator of double-stranded DNA-
			mediated type I interferon induction where it acts as an
			interferon-inducible E3 ubiquitin ligase that modulates
			STING to confer double-stranded DNA-mediated
32956	TRIM56	21074459	innate immune responses.
			TRIM56 is an anti-viral host factor that restricts
32956	TRIM56	21289118	pestivirus infection, specifically it was found to impair
52930	I KIIVI30	21289118	bovine viral diarrhoea virus replication. TRIM56 is an essential component of the TLR3 anti-
32956	TRIM56	22948160	viral signalling pathway.
			TRIM56 was identified in a systematic screen for
32956	TRIM56	23438823	positive regulators of innate immune responses.
			TRIM50 was identified in a systematic screen for
19813	TRIM50	23438823	positive regulators of innate immune responses.
24165	TD IX (1.2	22420022	TRIM13 was identified in a systematic screen for
34165	TRIM13	23438823	positive regulators of innate immune responses. TRIM9 was identified in a systematic screen for
6467	TRIM9	23438823	positive regulators of innate immune responses.
0107	11(11)1)	23 13 00 23	TRIM8 was identified in a systematic screen for
88134	TRIM8	23438823	positive regulators of innate immune responses.
			TRIM71 was identified in a systematic screen for
23894	TRIM71	23438823	positive regulators of innate immune responses.
107002		22428822	TRIM58 was identified in a systematic screen for
107983	TRIM58	23438823	positive regulators of innate immune responses.
69064	TRIM47	23438823	TRIM47 was identified in a systematic screen for positive regulators of innate immune responses.
02001		23 13 00 23	TRIM32 targets TMEM173 for ubiquitination and
			enhances the induction of IFNB against RNA and DNA
82716	TRIM32	22745133	viruses.
			TRIM32 was identified in a systematic screen for
82716	TRIM32	23438823	positive regulators of innate immune responses.

300948 24923	TRIM26 TRIM21	23438823 20627395	TRIM26 was identified in a systematic screen for positive regulators of innate immune responses. TRIM21 (Ro52) conjugates phosphorylated IKBKB with monoubiquitin and the IKBKB-induced NFkappa B signalling is downregulated. TRIM21-mediated monoubiquitination is involved in the subcellular translocation of active IKBKB to autophagosomes. TRIM21 is a member of the TRIM family of single-
24923	TRIM21	20668674	protein E3 ligases and TRIM21-mediated ubiquitination promotes the degradation of IRF7 following TLR7 and TLR9 stimulation. As TRIM21 is also IFN-inducible, this system constitutes a negative- feedback loop that acts to protect the host from the prolonged activation of the immune response. Tyrosine phosphorylation of the E3 ubiquitin ligase
24923	TRIM21	22479513	TRIM21 positively regulates interaction with IRF3 and hence TRIM21 activity.
24923	TRIM21	23438823	TRIM21 was identified in a systematic screen for positive regulators of innate immune responses.
43046	MID1	23438823	MID1 was identified in a systematic screen for positive regulators of innate immune responses.
75795	TRIM15	23077300	TRIM15 is required for RIG-I mediated interferon production to inhibit viral replication.
75795	TRIM15	23438823	TRIM15 was identified in a systematic screen for positive regulators of innate immune responses.
62871	TRIM7	23438823	TRIM7 was identified in a systematic screen for positive regulators of innate immune responses. STAT2 is part of STAT-containing transcription factor,
40906	STAT2	8943351	the alpha-interferon-induced ISGF3, that is composed of a STAT1:2 heterodimer and a weak DNA-binding protein, IRF9. STAT2 is a critical transactivator component of the interferon-stimulated gene factor 3 (ISGF3) complex
40906	STAT2	16689942	<ul><li>that drives the expression of many interferon (IFN)-inducible genes.</li><li>A loss-of-function mutation in STAT2 is associated</li></ul>
40906	STAT2	23391734	with increased susceptibility to childhood viral diseases.
64053	IL1R2	23395675	IL1R2 binds to IL1A to limit post-necrotic inflammation.
92888	MUL1	23399697	MUL1 limits inflammation by regulating the RIG-I anti-viral response. Creb1 over-expression in myeloid cells results in
161359	Creb1	23405224	<ul><li>increased abscess formation and aberrant cytokine response.</li><li>Lgals9 attenuates acute lung injury by preferentially suppressing pro-inflammatory functions in pDC-like</li></ul>
203274	Lgals9	21562126	macrophages through TLR2/TLR4 down-regulation.
203274	Lgals9	23408620	LGALS9 engagement impairs the cytotoxicity function and cytokine production of natural killer cells.

193040	Gabarap	23427251	Gabarap deficient mice are more susceptible to sepsis due to enhanced inflammasome activation. S100A9 is an intracellular calcium-binding protein that
102644	S100A9	12626582	promotes neutrophil/monocyte recruitment at inflamed tissues by enhancing attachment to endothelial cells. S100A9 is a myeloid-related protein that rapidly modulates more phase ritric avide production during
102644	S100A9	18714033	<ul> <li>modulates macrophage nitric oxide production during innate immune response.</li> <li>S100A9 (calgranulin B) and S100A8 (calgranulin A) form an antimicrobial heterodimeric complex known as calprotectin. Bacterial flagellin induces the upregulation of S100A9/S100A8 heterodimer via a TLR5-dependent mechanism in epidermal</li> </ul>
102644	S100A9	20555353	keratinocytes. S100A9 forms a complex with S100A8 and the complex is the site of interplay between extracellular Ca(2+) entry and intra-phagosomal reactive oxygen species production. S100A8 :: S100A9 acts as Ca(2+) sensor in phagosomal ROS production.
102644	S100A9	21239714	
102644	S100A9	21382888	S100A9-deficient murine neutrophils exhibited a reduce secretion of cytokines in response to TLR4 stimulation. In contrast, S100A9-deficient dendritic cells showed an exacerbated release of cytokines after TLR stimulation. S100A9 has no effect on the inflammatory status of macrophages. (Demonstrated in murine model)
			S100A9 is strongly upregulated in neutrophils upon bacterial infection, and sequesters zinc as a mechanism of nutritional immunity. Salmonella typhimurium overcomes this defence mechanism by expressing a
102644	S100A9	22423963	high affinity zinc transporter. (Demonstrated in mice) S100A9 forms a heterodimer with S100A8 and is a key player in protective innate immunity during Klebsiella
102644	S100A9	23133376	pneumonia infection. S100A8::S100A9 heterodimer sequesters Mn(2+) and
102644	S100A9	23431180	Zn(2+) to starve bacteria of these essential nutrients. MIRLET7B post-transcriptionally suppresses TLR4
126513	MIRLET7B	23437218	and regulates NF-kB-mediated responses in H. pylori infection. IL1R1 upregulates Mir135b as a negative feedback
221470	Mir135b	23440414	regulatory mechanism to resolve cigarette smoke- induced inflammation in the lung. Tlr1 :: Tlr2 dimeric pairs recognize malarial glycosylphosphatidylinositols (GPI) to initiates
165889	Tlr1	21439957	intracellular signalling and the production of pro- inflammatory cytokines.
165889	Tlr1	22778390	Tlr1 is a critical innate receptor for protective intestinal T(H)17 immunity against Yersinia enterocolitica. TLR1 in intestinal epithelium mediates activation and
165889	Tlr1	23443468	recruitment of dendritic cells to mount mucosal immunity against Yersinia enterocolitica.

			LEP upregulates TLR2 expression in monocytes, and may potentiate innate immunity and inflammation in hyperleptinemia conditions, such as obesity and type 2
39587	LEP	23341537	diabetes mellitus.
39587	LEP	23341537	LEP (leptin) upregulates TLR2 in human monocytes. Caspase-11 (SCAF11) increases susceptibility to
178155	Scaf11	22895188	Salmonella infection in the absence of caspase-1. SCAF11 (Caspase 11) is required for innate
178155	Scafl 1	23348507	surveillance against bacteria that escape the vacuole. Naip5 regulates the specificity of the Nlrc4 inflammasome for distinct bacterial ligands such as flagelllin.
177193	Naip5	21282416	
			Naip5 is a NOD-like receptor protein that directly and specifically binds to bacterial flagellin. Naip5 is a universal component of the flagellin-Nlrc4
177193	Naip5	21918512	inflammasome pathway. Microglial cells employ the NAIP5/NLRC4
177193	Naip5	23355222	inflammasome to monitor and clear central nervous system infections by flagellated bacteria.
194638	Fer	23355730	FER is an inhibitory kinase for neutrophil chemotaxis.
			RAD23A is a negative regulator of the RIG-I anti-viral signalling pathway by mediating the ubiquitin-
174863	Rad23a	23357418	dependent proteasomal degradation of TRAF2. RAD23A is a negative regulator of the RIG-I anti-viral
31509	RAD23A	23357418	signalling pathway by mediating the ubiquitin- dependent proteasomal degradation of TRAF2. Lum facilitates early innate immune responses and orchestrates bacterial clearance and inflammation
187527	Lum	23358433	resolution in the cornea. Agonists of TACR1 promote immunostimulatory dendritic cells by inhibiting IL10 synthesis and
158953	Tacr1	23365459	secretion. The bile acid sensor Nr1h4 (FXR) Is required for
184281	Nr1h4	23372731	immune-regulatory activities of TLR9 in intestinal inflammation.
			Siglecg is a member of the SIGLEC protein family that recognize sialoside-based patterns and responds selectively to danger associated molecular patterns (DAMPs) to initiate limited innate response.
176715	Siglecg	21208791	
176715	Siglecg	23374343	Siglecg expression is upregulated by RNA viruses to suppress the RIG-I anti-viral signalling pathway. CNOT8 negatively regulates interferon IFN/STAT1
55066	CNOT8	23386060	signalling pathway and its depletion results in enhanced response to viral infection.
147704	Chat	23297238	Chat is expressed by lymphocytes to regulate local recruitment of neutrophils.
62465	COX5B	23308066	COX5B coordinates with the autophagy pathway to control MAVS aggregation and antiviral signalling.
158216	Abl1	23325923	ABL1 kinase is required for neutrophil migration.

89599	ABL1	23325923	ABL1 kinase is required for neutrophil migration.
142854	Prkcd	22265677	Prkcd is an essential signalling kinase in C-type lectin receptor-mediated innate immunity and host protection.
142854	Prkcd	23326354	Suppression of Prkcd expression inhibited nitric oxide production in activated macrophages. Tbx21 (T-bet) is a member of the T-box transcription factor family that plays a critical role in interferon (UED) common regulation and promotion of call
210153	Tbx21	11752460	<ul><li>(IFN)-gamma regulation and promotion of cell- mediated immune responses against intracellular pathogens.</li><li>Tbx21 (T-bet) regulates the interplay between mucosal dendritic cells, innate lymphoid cells, and the intestinal</li></ul>
210153	Tbx21	23063332	microbiota.
210153	Tbx21	23334414	Gradual expression of Tbx21 controls the fate and function of innate lymphoid cells.
223707	Mir212	23264652	Mir212 targets Irak4 to dampen the inflammatory response in LPS-stimulated monocytes.
126933	MIR212	23264652	MIR212 targets IRAK4 to dampen the inflammatory response in LPS-stimulated monocytes.
221348	Mir132	23264652	Mir132 targets Irak4 to dampen the inflammatory response in LPS-stimulated monocytes. Hsa-mir-132 regulates antiviral innate immunity through suppression of the EP300 transcriptional co- activator, rather than a transcription factor or signaling protein. Furthermore, EP200, regulates has mir 122
127301	hsa-mir-132	20418869	protein. Furthermore, EP300 regulates hsa-mir-132 levels, revealing a dynamic equilibrium between hsa- mir-132 and EP300.
127301	hsa-mir-132	22822053	MIR132 is upregulated in lung cells in response to infection with influenza A virus.
127301	hsa-mir-132	23264652	MIR132 targets IRAK4 to dampen the inflammatory response in LPS-stimulated monocytes.
164070	Cryab	23242137	CRYAB is a suppressor of neuroinflammation, and the suppression is mediated by DRD2 in astrocytes. DRD2 activation in astrocytes suppresses
162803	Drd2	23242137	neuroinflammation in the central nervous system through CRYAB. Tph1-deficient mice lack non-neuronal serotonin, and
186197	Tph1	23243271	show impairment in neutrophil recruitment to sites of acute inflammation.
547297	ITGB3	23150579	ITGAV::ITGB3 is a sensor and activator of innate immunity to herpes simplex virus-1.
77054	ITGAV	23150579	ITGAV::ITGB3 is a sensor and activator of innate immunity to herpes simplex virus-1. Fcnb is crucial for the induction of innate immunity
151717	Fcnb	23150716	against pneumococcal infection through the lectin complement pathway.

			Fcna interact with mannose-binding lectin and fibrinogen/fibrin to augment the lectin complement pathway suggesting that this pathway collaborates with the coagulation system in the first-line host defence
146780	Fcna	20375621	against pathogens under conditions such as injury and inflammation. Fena is crucial for the induction of innate immunity
146780	Fcna	23150716	against pneumococcal infection through the lectin complement pathway. Glrx deficiency impairs neutrophil polarization,
168209	Glrx	23159440	chemotaxis, adhesion, and phagocytosis. HSPA1B is released extracellularly upon measles virus
606687	Hspa1b	23135720	infection, and induces Ifnb transcription in microglial cells. MIR497 inhibits IL1-induced IL6 transcription by
221234	Mir497	23092882	targeting MAPK/ERK pathway. MOV10 associates with LINE1 ribonucleoprotein
101151	MOV10	23093941	particles and restricts endogenous retrovirus transposition. DAGLB is a key metabolic hub within a lipid network
207742	Daglb	23103940	that regulates proinflammatory responses in peritoneal macrophages. Antiviral stress granules containing Ddx58 (RIG-I) and
199014	Eif2ak2	22912779	Eif2ak2 (PKR) have a critical role in viral detection and innate immunity.
199014	Eif2ak2	23115300	Herpes simplex virus evades antiviral host defence by interacting with EIF2AK2 (PKR) to inhibit autophagy. AKT1 is a Ser/Thr protein kinase that plays a pivotal role in functional activation in macrophages. AKT1 specifically functions in phagocytosis, intracellular bacterial infection, LPS tolerance, production of
22709	AKT1	21196185	inflammatory cytokines/mediators, and migration during macrophage-mediate innate immunity. AKT1 activation is blocked by Bacillus anthracis, resulting in the opening of a connexin ATP release channel and induction of macrophage death. Constitutive activation of AKT1 interferes with inflammasome activation and IL1B production, which
22709	AKT1	21683629	compromises antimicrobial immunity. (Demonstrated in murine model) AKT1 functions downstream of TLR2-stimulation to
22709	AKT1	22218715	induce the expression of the monocyte chemoattractant protein 1, CCL2. (Demonstrated in mice) AKT1 is important for establishing the inflammatory
22709	AKT1	23060458	microenvironment in the airway upon exposure to rhinovirus. AXL is a pleiotropic inhibitor of the innate immune response in dendritic cells, this tyrosine protein kinase is induced by IFNAR/STAT1 signalling and TLR ligation predominantly through TLR activation of the
52651	AXL	18083102	feed-forward cytokine pathway

			AXL regulates survival and migration of human
52651	AXL	19657094	dendritic cells by an IFN-alpha-inducible AXL/GAS6 pathway.
			TGF-Î <sup>2</sup> 1-induced AXL enhances apoptotic cell uptake and blocks proinflammatory cytokine production in the
52651	AXL	23071254	skin.
			MIR187 directly targets TNFA mRNA stability and translation and acts as an effector of IL10-downstream
127585	MIR187	23071313	signalling to inhibit pro-inflammatory cytokines. Pparg negatively regulates Ifnb production in Tlr3/4 stimulated macrophages by preventing Irf3 binding to
			the Ifnb promoter.
179253	Pparg	21148557	
			The loss of PPAR $\hat{I}^3$ in T cells increased colitis disease activity and colonic inflammatory lesions following
179253	Pparg	23071818	Clostridium difficile infection.
			TRIM62 is required for TRIF-mediated NFkB, AP-1
95888	TRIM62	23077300	and interferon production after LPS challenge in macrophages.
20000	11(11)(02	23077300	Mir4661 directly targets Ifna expression to inhibit host
272448	Mir4661	23042536	antiviral immune responses.
			Pyhina is an activator of the STING-dependent IFN and ASC-dependent inflammasome in antiviral innate
205966	AI607873	23045604	immunity.
			Pydc3 is an activator of the STING-dependent IFN and
205951	Pydc3	23045604	ASC-dependent inflammasome in antiviral innate immunity.
			Pyhin1 is an activator of the STING-dependent IFN
205927	Dybin 1	23045604	and ASC-dependent inflammasome in antiviral innate
203927	Pyhin1	23043004	immunity. Pyhinb is an activator of the STING-dependent IFN
			and ASC-dependent inflammasome in antiviral innate
205894	BC094916	23045604	immunity. Pyblhinc is an activator of the STING-dependent IFN
			and ASC-dependent inflammasome in antiviral innate
205885	Gm4955	23045604	immunity.
			TRIM28 interacts with IRF5 and mediates heterochromatinization at M1 macrophage markers
73191	TRIM28	22995936	(e.g. TNF) to suppress transcription.
			CALCOCO2 recognizes ubiquitin-coated Salmonella
			enterica in human cells and binds to the adaptor proteins AZI2 and TBKBP1 to recruit TBK1.
			CALCOCO2 also recruits LC3, an autophagosomal
57242		10020700	marker, to activate autophagy against bacteria
57242	CALCOCO2	19820708	attempting to colonize their cytosol. CALCOCO2 is a novel receptor for the selective
			autophagy of cytosolic bacteria. CALCOCO2 (NDP52)
			binds to the bacterial ubiquitin coat as well as to ATG8/LC3 and delivers cytosolic bacteria into
57242	CALCOCO2	20104023	ATG8/LC3 and delivers cytosolic bacteria into autophagosomes.

57242	CALCOCO2	19841643	CALCOCO2 directly binds to ubiquitinated bacteria and facilitates the assembly of an autophagic membrane that surrounds bacterial invaders. CALCOCO2 and SQSTM1 are ubiquitin-autophagy receptors that are required for the recognition of extracelluar bacterial DNA by the TMEM173 (STING)-dependent cytosolic pathway, marking bacteria with ubiquitin and delivery of bagilli to
57242	CALCOCO2	22901810	bacteria with ubiquitin, and delivery of bacilli to autophagosomes. (Demonstrated in mouse) MAP1LC3C (LC3C) and its receptor CALCOCO2
57242	CALCOCO2	23022382	(NDP52) are essential for antibacterial autophagy. MAP1LC3C (LC3C) and its receptor CALCOCO2
107725	MAP1LC3C	23022382	(NDP52) are essential for antibacterial autophagy. Foxo3 is an Ikbke-controlled checkpoint of IRF activation and regulation of Ifnb expression.
147382	Foxo3	22531926	(Demonstrated in human) FOXO3 is a negative regulator of Irf7 transcription,
147382	Foxo3	22982991	and forms a coherent feed-forward regulatory circuit with Irf7 and Ifnb to maximize antiviral responses. FOXO3 is a transcriptional activator which targets genes FAS ligand and TNFSF10 (TRAIL), involved in the extrinsic apoptotic pathway, and BBC3 (PUMA),
95024	FOXO3	19050264	PMAIP1 (Noxa), and BCL2L11 (Bim), which are part of the intrinsic apoptotic pathway. FOXO3, along with AKT, is involved in TLR9- mediated anti grantesis and is a distinct membrane for
95024	FOXO3	20739833	mediated anti-apoptosis and is a distinct regulator for FLICE-like inhibitory protein (FLIP) expression. FOXO3 is an IKBKE-controlled checkpoint of IRF
95024	FOXO3	22531926	activation and regulation of IFNB expression. OXO3 is a negative regulator of IRF7 transcription, and forms a coherent feed-forward regulatory circuit with IRF7 and IFNB to maximize antiviral responses.
95024	FOXO3	22982991	(Demonstrated in mice) VTRNA2-1 attenuates EIF2AK2 (PKR) activation, and it may act as a negative regulator of innate immune
115355	VTRNA2-1	22986343	response to dsRNA viruses. ADIPOQ can induce pro-inflammatory functions in
149533	Adipoq	22948153	ADIPOQ binds to C1Q and activates the classical
69167	ADIPOQ	18179772	pathway of complement. ADIPOQ can induce pro-inflammatory functions in
69167	ADIPOQ	22948153	macrophages and T cells. Trim56 is an anti-viral host factor that restricts pestivirus infection, specifically it was found to impair
204815	Trim56	21289118	bovine viral diarrhoea virus replication. TRIM56 is an essential component of the TLR3 anti-
204815	Trim56	22948160	viral signalling pathway. (Demonstrated in human) Tnfaip8l2 is a negative regulator of immunity that controls innate immunity to RNA by targeting the
171590	Tnfaip8l2	22904303	PI3K-Rac pathway. Tnfaip812 (TIPE2) serves as a negative regulator of
171590	Tnfaip8l2	22949657	phagocytosis and oxidative burst during infection.

			TNFAIP8L2 is a negative regulator of innate and adaptive immunity that maintains immune homeostasis
102300	TNFAIP8L2	18455983	by binding to CASP8 and inhibiting nuclear factor kappa B activation while promoting apoptosis. TNFAIP8L2 is a negative regulator of immunity that
102300	TNFAIP8L2	22904303	controls innate immunity to RNA by targeting the PI3K-Rac pathway. (Demonstrated in mouse) TNFAIP8L2 (TIPE2) serves as a negative regulator of
102300	TNFAIP8L2	22949657	<ul><li>phagocytosis and oxidative burst during infection.</li><li>(Demonstrated in mice)</li><li>Abca1 deletion in the myeloid lineage enhances host</li></ul>
147642	Abcal	22955730	immune response and clearance of Listeria monocytogenes. ABCA1 promotes the efflux of bacterial
79441	ABCA1	20472936	lipopolysaccharide (LPS) from macrophages and accelerates recovery from LPS-induced tolerance. ABCA1 in macrophages dampens inflammation by reducing MYD88-dependent TLR trafficking to lipid
79441	ABCA1	20650929	rafts, thus selectively reducing free cholesterol content in lipid rafts. ABCA1 deletion in the myeloid lineage enhances host
79441	ABCA1	22955730	immune response and clearance of Listeria monocytogenes. (Demonstrated in mice) Mir467b regulates lipid accumulation and
224167	Mir467b	22963823	proinflammatory cytokine secretion in macrophages by targeting the LPL gene. Trp73 is required for macrophage-mediated innate
206900	Trp73	22976836	immunity and the resolution of inflammatory responses. TP73 is required for macrophage-mediated innate
86757	TP73	22976836	immunity and the resolution of inflammatory responses. (Demonstrated in mouse) MIR10B directly targets the 3'UTR of MICB and
127521	MIR10B	22915757	down-regulates its expression, which may aid tumour cells in immune escape from natural killer cells. IL1RL1 acts as a negative regulator of TLR2 signalling whereby over-expression of IL1RL1 can dose- dependently attenuate bacterial lipoprotein (BLP)-
64166	IL1RL1	20400705	<ul><li>induced NF-kappaB activation, but is not required for BLP-induced tolerance.</li><li>IL1RL1 is the receptor for IL33, a factor shown to modulate tryptase expression in mesenchymal cells (MCs), identifying a novel pathway by which MCs exposed to inflammatory cytokines modulate the</li></ul>
64166	IL1RL1	20427273	phenotype of local MCs to shape their immune responses. IL1RL1 (ST2) pre-treatment suppresses cytokine
64166	IL1RL1	22922442	production and inhibits LPS signalling in dendritic cells.

			IL15 is a pluripotent anti-apoptotic cytokine that signals to cells of both the innate and adaptive immune system and is regarded as a highly promising immunomodulatory agent in cancer therapy. IL15 prevents two immunopathologic hallmarks of sepsis, namely, apoptosis and immunosuppression, and
39028	IL15	20026737	improves survival in two different models of sepsis. IL15 regulates homeostasis and terminal maturation of
39028	IL15	22084435	NKT cells. (Demonstrated in mice) IL15 secreted by inflammatory monocytes is critical for the differentiation of memory CD8(+) T and NK
39028	IL15	22940097	lymphocytes into antimicrobial effector cells. (Demonstrated in mice) IRF8 interacts with TRAF6 to modulate TLR
45278	IRF8	16484229	signalling and may also contribute to the cross-talk between IFN-gamma and TLR signalling pathways. IRF8 belongs to a family of interferon (IFN) regulatory factors that modulate various important physiologic
45278	IRF8	18469857	processes including host defence, cell growth and differentiation and immune regulation. De-SUMOylation of IRF8 at residue Lys310 acts as a molecular mechanism to trigger innate immune
45278	IRF8	22942423	responses in activated macrophages. (Demonstrated in mice) Calcoco2 and Sqstm1 are ubiquitin-autophagy receptors that are required for the recognition of
209835	Calcoco2	22901810	extracelluar bacterial DNA by the Tmem173 (STING)- dependent cytosolic pathway, marking bacteria with ubiquitin, and delivery of bacilli to autophagosomes. SQSTM1 (p62) targets invading bacteria to the autophagy pathway and its expression is required for efficient autophagy of bacteria, as well as restriction of
61811	SQSTM1	19812211	their intracellular replication. SQSTM1 is a key intracellular target of innate defence
61811	SQSTM1	19850933	regulator-1 (IDR-1), a synthetic peptide. SQSTM1 has bactericidal properties where it brings cytosolic proteins to autolysosomes where they are
61811	SQSTM1	20206555	processed from innocuous precursors into neo- antimicrobial peptides. SQSTM1 and HDAC6 are important determinants of aggregated localization of MyD88 and MyD88 activation initiates a polyubiquitinated protein
61811	SQSTM1	20837465	accumulating pathway that modulates MyD88- dependent signal transduction. SQSTM1 is required for TLR4-mediated autophagy. TLR4-driven induction of SQSTM1 plays an essential role in the formation and the autophagy degradation of aggresome-like induced structures, which might be
61811	SQSTM1	21220332	critical for regulating host defence.

			SQSTM1 and CALCOCO2 are ubiquitin-autophagy receptors that are required for the recognition of extracelluar bacterial DNA by the TMEM173 (STING)-dependent cytosolic pathway, marking
61811	SQSTM1	22901810	bacteria with ubiquitin, and delivery of bacilli to autophagosomes. (Demonstrated in mouse) The chymase mouse mast cell protease 4 (Mcpt4)
166604	Mcpt4	22901752	degrades TNF, limits inflammation, and promotes survival in a model of sepsis. Caspase-11 (SCAF11) increases susceptibility to Salmonella infection in the absence of caspase-1.
28456	SCAF11	22895188	(Demonstrated in mouse) Hyperactivated Ern1 (IRE11±) increases Txnip mRNA stability by reducing levels of a Txnip destabilizing microRNA, miR-17. In turn, elevated Txnip protein activates the Nlrp3 inflammasome, causing procaspase- 1 cleavage and interleukin $1 \hat{1}^2$ (IL-1 $\hat{1}^2$ ) secretion.
175493	Txnip	22883233	(Demonstrated in human) NLRC4 is part of the NOD-like receptor family of proteins that consists of more than 20 related family members, that is present in the cytosol and recognizes
43398	NLRC4	18487086	intracellular ligands. NLR4C and other NLR proteins function to control IL- 1, NF-kappaB, and host response to pathogens, including distinct forms of cell death. NLR4C is
43398	NLRC4	18280719	important for CASP1 activation and IL-1 processing. NLRC4 expressed in monocytes associates with NOD2 following exposure to bacterial peptidoglycan, implying a regulatory role for interaction of NACHT- domain containing proteins in the innate immune
43398	NLRC4	15107016	response. NLR4C detects the basal body rod component of the T3SS apparatus (rod protein) from S. typhimurium (PrgJ), Burkholderia pseudomallei (BsaK), Escherichia coli (EprJ and EscI), Shigella flexneri (MxiI), and Pseudomonas aeruginosa (PscI) and the specific detection of the virulence machinery permits the discrimination between pathogenic and non-pathogenic
43398	NLRC4	20133635	bacteria. The NLRC4 inflammasome is important for control of mucosal Candida infection, impacting on inflammatory cell recruitment to infected tissues, as well as protecting against the systemic dissemination of
43398	NLRC4	22174673	infection. (Demonstrated in mouse) Flagellin-induced NLRC4 inflammasome activation in splenic dendritic cells triggers antigen-independent
43398	NLRC4	22231517	IFN-gamma production by memory T cells. (Demonstrated in mouse) NLRC4 is important for host survival and bacterial clearance, as well as neutrophil-mediated inflammation
43398	NLRC4	22547706	in the lungs following Klebsiella pneumoniae infection. (Demonstrated in mouse)

			NLRC4-dependent production of IL1B by intestinal phagocytes is a mechanism that discriminates
43398	NLRC4	22484733	pathogenic from commensal bacteria in the intestinal host defence. (Demonstrated in mouse) NLRC4 Ser †533 phosphorylation is essential for
			procaspase-1 recruitment to the NLRC4 inflammasome complex after S. typhimurium infection. (Demonstrated
43398	NLRC4	22885697	in mouse) Il12, consisting of Il12a and Il12b subunits, induces
150825	Il12a	22888135	Il2ra to form high-affinity Il2 receptors on natural killer cells in response to mouse cytomegalovirus infection.
			Tlr13 recognizes a conserved 23S rRNA sequence that is the binding site of macrolide, lincosamide, and streptogramin group (MLS) antibiotics in bacteria. This reveals that specific mechanisms of antibiotic
166933	Tlr13	22821982	resistance are potent bacterial immune evasion strategies for avoiding recognition via Tlr13. Nlrc3 inhibits Toll-like receptor (TLR)-dependent activation of NF-kB by interacting with the TLR
129350	Nlrc3	22863753	signalling adaptor Traf6 to attenuate ubiquitination of Traf6 and activation of NF-kB.
			NLRC3 inhibits Toll-like receptor (TLR)-dependent activation of NF-kB by interacting with the TLR signalling adaptor TRAF6 to attenuate ubiquitination
12133	NLRC3	22863753	of TRAF6 and activation of NF-kB. (Demonstrated in mouse) MIR378 is specifically induced by IL4 via the IL-4-
126947	MIR378	22855601	Receptor/PI3K/AKT signalling pathway to drive macrophage proliferation. (Demonstrated in mouse ) Cftr deficiency alters the innate immunity of the biliary epithelium and reduces endotoxin tolerance, resulting
129229	Cftr	21712022	in increased inflammatory response mediated by Tlr4 and NFkB.
			Cftr is involved in myeloid cell function and its absence from myeloid-derived cells slows resolution of
129229	Cftr	22859830	inflammation and infection of the lung. CFTR deficiency alters the innate immunity of the
			biliary epithelium and reduces endotoxin tolerance, resulting in increased inflammatory response mediated
37713	CFTR	21712022	by TLR4 and NFkB. CFTR is involved in myeloid cell function and its
			absence from myeloid-derived cells slows resolution of inflammation and infection of the lung. (Demonstrated
37713	CFTR	22859830	in mouse) Mmp9 and Neu1 cross-talk in alliance with Tlr4 on the
			cell surface is a novel membrane sialidase-controlling mechanism that depends on ligand binding to its Toll- like receptor (TLR) to induce Neu1 activity, to influence receptor desialylation and subsequently to induce TLR receptor activation and the production of
212451	Mmp9	21873432	nitric oxide and pro-inflammatory cytokines in dendritic and macrophage cells.

212451	Mmp9	22496659	Mmp9 inhibits influenza virus pathogenesis by mediating neutrophil migration to the respiratory tract.
212451	Mmp9	22860023	Mmp9 cleaves the pulmonary collectin Sftpd (SP-D) leading to loss of its innate immune functions. MMP9 and NEU1 cross-talk in alliance with TLR4 on the cell surface is a novel membrane sialidase- controlling mechanism that depends on ligand binding to its Toll-like receptor (TLR) to induce NEU1 activity, to influence receptor desialylation and subsequently to induce TLR receptor activation and the production of nitric oxide and pro-inflammatory cytokines in dendritic and macrophage cells. (Demonstrated in
78722	MMP9	21873432	murine model) MMP9 inhibits influenza virus pathogenesis by mediating neutrophil migration to the respiratory tract.
78722	MMP9	22496659	(Demonstrated in mice) MMP9 cleaves the pulmonary collectin SFTPD (SP-D) leading to loss of its innate immune functions.
78722	MMP9	22860023	(Demonstrated in mouse) Ahsg is an endogenous ligand of Tlr4 that promotes
148706	Ahsg	22842477	inflammatory signalling leading to lipid-induced insulin resistance. AHSG is an endogenous ligand of TLR4 that promotes
68916	AHSG	22842477	inflammatory signalling leading to lipid-induced insulin resistance. (Demonstrated in mice) Map3k7 activation is impaired during endotoxin tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of anti-inflammatory or anti- microbial mediators.
132962	Map3k7	21220427	Map3k7 polyubiquitination is essential for the activation of NF-kB signalling downstream of TNF
132962	Map3k7	22069318	receptor, IL1 receptor and Tlr4. Map3k7 is necessary for the neutrophil priming effect
132962	Map3k7	22843747	of leukotriene B (4) to enhance TLR stimulation. Olfm4 is a negative regulator of neutrophil bactericidal activity by restricting cathepsin C-mediated protease
183156	Olfm4	22844115	activities. OLFM4 exerts considerable influence on the host defence against H. pylori infection acting through NOD1 and NOD2 mediated NF-kappaB activation and subsequent cytokines and chemokines production, which in turn inhibit host immune response and
36123	OLFM4	20534456	contribute to persistence of H. pylori colonization. OLFM4 is a negative regulator of neutrophil bactericidal activity by restricting cathepsin C-
36123	OLFM4	22844115	mediated protease activities. (Demonstrated in mice)

			Itgam :: Itgb2 is the principal leukocyte receptor involved in the recognition of the fungus Candida albicans. Recognition of Pra1p protein of C. albicans by Itgam :: Itgb2 plays a pivotal role in determining fungal virulence, and host response/protection against C. albicans infection.
167243	Itgb2	21245270	Itgb2 is involved in cell-cell contact signalling between activated apoptotic lymphocytes and dendritic cells
167243	Itgb2	22396536	<ul><li>(DC) during the maturation of DCs. (Demonstrated in human)</li><li>Itgax::Itgb2 is a leukocyte receptor for Candida</li></ul>
167243	Itgb2	22844116	albicans and is essential for protection against fungal infections. ITGAM :: ITGB2 is the principal leukocyte receptor involved in the recognition of the fungus Candida albicans. Recognition of Pra1p protein of C. albicans by ITGAM :: ITGB2 plays a pivotal role in determining fungal virulence, and host response/protection against C. albicans infection. (Demonstrated in murine model)
5590	ITGB2	21245270	ITGB2 is involved in cell-cell contact signalling between activated apoptotic lymphocytes and dendritic
5590	ITGB2	22396536	cells (DC) during the maturation of DCs. ITGAX::ITGB2 is a leukocyte receptor for Candida albicans and is essential for protection against fungal
5590	ITGB2	22844116	infections. Itgax is the main marker for identification of dendritic cells (DCs). Itgax expression negatively orchestrates both adaptive and innate immunity against herpes simplex virus type 1 (HSV-1) ocular infection via higher activities of type 1 interferon and CD8(+) T cell
210550	Itgax	21775452	responses. Itgax::Itgb2 is a leukocyte receptor for Candida albicans and is essential for protection against fungal
210550	Itgax	22844116	infections. ITAGX is the main marker for identification of dendritic cells (DCs). ITGAX expression negatively orchestrates both adaptive and innate immunity against herpes simplex virus type 1 (HSV-1) ocular infection via higher activities of type 1 interferon and CD8(+) T
28212	ITGAX	21775452	cell responses. (Demonstrated in murine model) ITGAX::ITGB2 is a leukocyte receptor for Candida albicans and is essential for protection against fungal
28212	ITGAX	22844116	infections. Ikbke and other IKK kinases regulate each other by an intricate network involving phosphorylation of their catalytic and regulatory (NEMO, TANK) subunits to balance their activities during innate immunity.
190983	Ikbke	21138416	balance then activities during innate ininumity.

			Ikbke kinase modulates II-17 signalling through Traf3ip2 adaptor protein phosphorylation, resulting in the induction of neutrophilia and pulmonary
190983	Ikbke	21822257	inflammation. Ikbke is an important signalling molecule in the IFN
190983	Ikbke	22065572	pathway and is necessary to mount anti-viral immunity against West Nile virus. Ikbke plays a critical role in regulating the balance
190983	Ikbke	22171011	between the type I and type II interferon (IFN) signalling pathways. Ikbke is sequestered by arenavirus nucleoproteins to
190983	Ikbke	22532683	block its autocatalytic activity and its ability to active Irf3. (Demonstrated in human) Masp2 deficient mice are defective in the lectin
204703	Masp2	22792067	pathway of complement activation and are highly susceptible to pneumococcal infection. Ms4a8a expression is induced by TLR signalling in
145846	Ms4a8a	22806454	M2-like macrophages. Ms4a8a is expressed at the late stages of Trypanosoma congolense and Taenia crassiceps infections. MS4A8B expression is induced by TLR signalling in M2-like macrophages. MS4A8B is expressed at the late
49293	MS4A8B	22806454	stages of Trypanosoma congolense and Taenia crassiceps infections. (Demonstrated in mice) Host activation of Casp7 in response to pore formation during Listeria monocytogene infection represents an
178813	Casp7	22807671	adaptive mechanism by which host cells can protect membrane integrity during infection. CASP7 is a substrate of the CASP1 inflammasomes, demonstrating the existence of a nucleotide binding and oligomerization domain-like receptor/CASP1/CASP7 cascade and the existence of distinct activation mechanisms for CASP3 and CASP7
90066	CASP7	18667412	in response to microbial stimuli and bacterial infection. CASP7 activation by the NLRC4 inflammasome
90066	CASP7	19343209	restricts Legionella pneumophila infection in mice. Host activation of CASP7 in response to pore formation during Listeria monocytogene infection represents an adaptive mechanism by which host cells can protect membrane integrity during infection.
90066	CASP7	22807671	(Demonstrated in mice)
281264	MIR1275	22822053	MIR1275 is upregulated in lung cells in response to infection with influenza A virus. MIR200C is upregulated in lung cells in response to
127093	MIR200C	22822053	infection with influenza A virus.
126579	MIRLET7C	22835429	MIRLET7C down-regulates IL10 expression. Pura expression is repressed by endogenous miRNA to restrict HIV-1 infection in monocytes. (Demonstrated
137597	Pura	22835829	in human) PURA expression is repressed by endogenous miRNA
48322	PURA	22835829	to restrict HIV-1 infection in monocytes.

184519	Ace2	22837003	Ace2 deficiency results in increased susceptibility to intestinal inflammation induced by epithelial damage. Ace2 links amino acid malnutrition to altered gut microbiota, which leads to colitis susceptibility. ACE2 deficiency results in increased susceptibility to intestinal inflammation induced by epithelial damage.
46534	ACE2	22837003	ACE2 links amino acid malnutrition to altered gut microbiota, which leads to colitis susceptibility. (Demonstrated in mice) The regulated phosphorylation of Eif4e has a key role in antiviral host defense by selectively controlling the translation of Nfkbia mRNA, which encodes a critical
196687	Eif4e	22544393	suppressor of the innate antiviral response. As a result, Eif4e null mice exhibit lower susceptibility to viral infections. The regulated phosphorylation of EIF4E has a key role in antiviral host defense by selectively controlling the translation of NFKBIA mRNA, which encodes a critical suppressor of the innate antiviral response. As a result, EIF4E null mice exhibit lower susceptibility to
20070		22544202	viral infections. (Demonstrated in mice)
30970	EIF4E	22544393	The leukocyte integrin antagonist (Edil3) Del-1 inhibits
170705	Edil3	22447028	IL-17-mediated inflammatory bone loss. The leukocyte integrin antagonist (EDIL3) Del-1
32547	EDIL3	22447028	inhibits IL-17-mediated inflammatory bone loss. (Demonstrated in mice)
2147	MAPK1	22447027	MAPK1 (ERK) and MAPK14 (p38) control the dynamic balance regulating neutrophil migration. NLRP4 binds with NLRC4 through NACHT domain, which by mediating hetero-oligomerization, creates protein-interaction networks that potentially modulate
71189	NLRP4	15107016	immune responses to invading pathogens.
71189	NLRP4	22388039	NLRP4 regulates the activation of type I interferon triggered by dsRNA or dsDNA. NF-ΰB-mediated degradation of the coactivator NRIP1 (RIP140) regulates inflammatory responses and contributes to endotoxin tolerance. (Demonstrated in
788	NRIP1	22388040	mice)
			NRIP1 is degraded by the NF-kB pathway to inactivate inflammatory gene expression and promotes endotoxin
788	NRIP1	22388040	tolerance. (Demonstrated in mice) Btk interacts with intracellular MHC class II molecules
171427	Btk	21441935	to activate adaptor molecules Myd88 and Ticam1 to promote TLR signalling. Btk is a positive regulator in the ITAM-mediated Trem1/Tyrobp pathway, which induces pro-
171427	Btk	21659545	inflammatory cytokines such as TNF-alpha, Il8, and activation/differentiation cell surface markers.

			Btk directly phosphorylates Tlr3 and plays a critical role in the induction of inflammatory cytokines and
171427	Btk	22454496	Ifnb.
171427	Btk	22589540	Btk is required for the activation of natural killer cells. Btk is a negative regulator of TLR- or TNF-stimulated reactive oxygen species (ROS) production in
171427	Btk	22366891	neutrophils. (Demonstrated in human) Zc3h12a prevents autoimmunity by controlling the stability of cytokine mRNA, such as Il6. Zc3h12a protein degradation is facilitated by IKK complex,
187492	Zc3h12a	22037600	which acts downstream of TLR stimulation. Zc3h12a is a potent regulator of innate immunity, which can be strongly engaged in the pathogenesis of acute and chronic infective diseases. (Demonstrated in
187492	Zc3h12a	22777400	human) ZC3H12A prevents autoimmunity by controlling the stability of cytokine mRNA, such as IL6. ZC3H12A protein degradation is facilitated by IKK complex,
96402	ZC3H12A	22037600	<ul><li>which acts downstream of TLR stimulation.</li><li>(Demonstrated in mice)</li><li>ZC3H12A is a potent regulator of innate immunity,</li><li>which can be strongly engaged in the pathogenesis of</li></ul>
96402	ZC3H12A	22777400	acute and chronic infective diseases. TLR1 polymorphisms affect innate immune responses
13627	TLR1	18635889	and outcomes in sepsis. TLR1 restrains potentially dangerous innate response
13627	TLR1	11932926	to LPS by binding to TLR4 and preventing the formation of active signalling complexes. TLR1 interacts with TLR2 to recognize the lipid
13627	TLR1	12077222	configuration of the native mycobacterial lipoprotein as well as several triacylated lipopeptides. TLR1 and TLR6 are involved in the discrimination of a
13627	TLR1	12697090	subtle difference between triacyl and diacyl lipopeptides through interaction with TLR2. TLR1 :: TLR2 dimeric pairs recognize malarial glycosylphosphatidylinositols (GPI) to initiates intracellular signalling and the production of pro-
13627	TLR1	21439957	inflammatory cytokines. TLR1 is a critical innate receptor for protective
13627	TLR1	22778390	intestinal T(H)17 immunity against Yersinia enterocolitica. (Demonstrated in mice) Deletion of P2ry14 inhibits macrophage recruitment
147059	P2ry14	22778393	and tissue inflammation, which mitigate diet-induced insulin resistance. Deletion of P2RY14 inhibits macrophage recruitment and tissue inflammation, which mitigate diet-induced
61501	P2RY14	22778393	insulin resistance. (Demonstrated in mice) Dusp10 protects against sepsis-induced acute lung
207813	Dusp10	22307906	injury. DUSP10 protects against sepsis-induced acute lung
106785	DUSP10	22307906	injury. (Demonstrated in mice)

			E2f1 is important for normal inflammatory response to systemic LPS by enhancing the production of Il6 and Tnfa in murine macrophages.
210416	E2f1	21131441	
210416	E2f1	22310660	E2f1 directly binds to the promoter of Tlr3 to inhibit transcription. E2F1 is important for normal inflammatory response to systemic LPS by enhancing the production of IL6 and TNFA in macrophages. (Demonstrated in murine model)
67214	E2F1	21131441	
67214	E2F1	22310660	E2F1 directly binds to the promoter of TLR3 to inhibit transcription. (Demonstrated in mice) Rb1 positively regulates Tlr3 expression by
179829	Rb1	22310660	modulating the transcription factor E2f1. RB1 positively regulates TLR3 expression by modulating the transcription factor E2F1.
32092	RB1	22310660	(Demonstrated in mice)
137812	Angpt1	22015631	Angpt1 promotes IL8 synthesis and release in neutrophils. (Demonstrated in human) ANGPT1 promotes IL8 synthesis and release in
32292	ANGPT1	22015631	neutrophils. Dok3 is a negative regulator of TLR signalling by limiting LPS-induced ERK activation and cytokine
156241	Dok3	22761938	responses in macrophages. DOK3 is a negative regulator of TLR signalling by
60370	DOK3	22761938	limiting LPS-induced ERK activation and cytokine responses in macrophages. (Demonstrated in mice)
168384	Dicer1	22252463	Down-regulation of Dicer1 elicits an interferon response in endometrial cancer cells. Down-regulation of DICER1 elicits an interferon
18522	DICER1	22252463	Down-regulation of DICER1 elicits an interferon response in endometrial cancer cells. Regulator of NF-kappaB and CASP1, angiotensin II
16041	NLRP6	12633874	nd vasopressin receptor NLRP6 is a negative regulator of inflammatory signalling and impedes the clearance of both Gram- positive and -negative bacterial pathogens.
16041	NLRP6	22763455	(Demonstrated in mice) Trim32 targets Tmem173 for ubiquitination and enhances the induction of Ifnb against RNA and DNA
156706	Trim32	22745133	viruses. (Demonstrated in human) Tufm inhibits RLR-induced IFN-I and promotes
209544	Tufm	22749352	autophagy during viral infection. (Demonstrated in human)
23469	TUFM	22749352	TUFM inhibits RLR-induced IFN-I and promotes autophagy during viral infection. NLRX1 is an amplifier of ROS generation and an inhibitor of antiviral mitochondrial signalling with markedly increased IFN- $\hat{1}^2$ responses in cells with
74346	NLRX1	18311173	down-regulated NLRX1 expression.

			NLRX1 depletion via siRNA promotes virus-induced
74346	NLRX1	18200010	type 1 interferon production and decreases viral replication. NLRX1 is an important regulator of the type I
74246	NI DV1	10200710	interferon (IFN) response, and prevents MAVS from signalling the activation of a type I IFN response by the
74346	NLRX1	18280719	RNA helicases NLRX1 deletion leads to constitutive interaction of MAVS and DDX58, resulting in increased inflammation. In addition to attenuating the antiviral
74346	NLRX1	21703540	response, NLRX1 directly associates with TRAF6 and inhibits NFkB pathway in LPS-activated macrophages. NLRX1 negatively regulates TLR-mediated NFkB activation by directly interacting with TRAF6 or IKK kinase. Nlrx1 knockdown in mice enhances their
74346	NLRX1	21703539	susceptibility to LPS-induced septic shock and increases plasma IL6 levels. NLRX1 attenuates type I IFN production and promotes
74346	NLRX1	22749352	autophagy during viral infection. mmu-mir-29a suppresses immune responses to intracellular pathogens by targeting IFN-gamma mRNA. Mice infected with Listeria monocytogenes or
221242	mmu-mir-29a	21785411	Mycobacterium bovis bacillus Calmette-Gu à © rin (BCG) downregulated miR-29 expression. mmu-mir-29a reduces the expression of IFN-alpha
221242	mmu-mir-29a	22179202	receptor and is critical for dampening the sensitivity of thymic epithelium to infection signals. Mir29a is secreted by tumour cells in the form of
221242	mmu-mir-29a	22753494	exosomes, and binds to intracellular Tlr7 in immune cells to trigger a prometastatic inflammatory response. Gnai2 is regulated by Tlr signaling and plays an anti- inflammatory role in endotoxema and polymicrobial sepsis.
198032	Gnai2	21255617	•
198032	Gnai2	22581266	Sustained activation of Gnai2 dampens Tnf and II6 production stimulated by Tlr2/4 ligands. GNAI2 is regulated by TLR signalling and plays an anti-inflammatory role in endotoxema and polymicrobial sepsis. (Demonstrated in murine model)
36105	GNAI2	21255617	Sustained activation of GNAI2 dampens TNF and IL6
36105	GNAI2	22581266	production stimulated by TLR2/4 ligands. (Demonstrated in mice) Spink5 binds to antimicrobial peptide dermcidin
144109	Spink5	22588119	(DCD) and is involved in the trafficking of DCD in the epidermis. (Demonstrated in human) SPINK5 binds to antimicrobial peptide dermcidin
52343	SPINK5	22588119	<ul><li>(DCD) and is involved in the trafficking of DCD in the epidermis.</li><li>Lst1 recruits tyrosine-phosphorylated Ptpn6 and</li></ul>
177285	Lst1	22589543	Ptpn11 to the plasma membrane in myeloid leukocytes. (Demonstrated in human)

			LST1 recruits tyrosine-phosphorylated PTPN6 and PTPN11 to the plasma membrane in myeloid
77937	LST1	22589543	leukocytes. Scarb1 regulates the macrophage inflammatory
201281	Scarb1	22589557	response to LPS.
63952	SCARB1	22589557	SCARB1 regulates the macrophage inflammatory response to LPS. (Demonstrated in mice) Spag11a is a beta-defensin expressed in epididymis.
274007	Spag11a	22535201	Overexpression of Spag11a confers resistance to E. coli infection in epididymis.
174941	Tsc22d3	22539300	Tsc22d3 is downregulated following TLR activation in macrophages.
81583	TSC22D3	22539300	TSC22D3 is downregulated following TLR activation in macrophages.
36000	NOS2	20157607	NOS2 is a nitric oxide synthase (iNOS) protein and combined treatment with 1,25-dihydroxyvitamin D3 (1,25-D3) and IFN-gamma has been shown to synergistically enhance NO synthesis and NOS2 expression induced by Mycobacterium tuberculosis (MTB) or by its purified protein derivatives in human monocyte-derived macrophages. NOS2 is an inducible nitric oxide (NO) synthase present in innate immune cells that produces NO in response to certain infections or upon stimulation with cytokines such as IFN-gamma and TNF. For optimal induction of NOS2 during Chlamydophila pneumoniae infection, the concerted action of the MyD88-
36000	NOS2	18799752	dependent transcription factors NF-kappaB and AP-1 and of the MyD88-independent transcription factors phosphorylated STAT1 and IRF1 is required. NOS2 (iNOS) and CALM1 (CaM) coordinately function to form a stable complex that is part of a rapid host response that functions within the first 30 min
36000	NOS2	16893173	following bacterial infection to up-regulate the innate immune system involving macrophage activation. Nitric oxide production by NOS2 promotes Listeria
36000	NOS2	22542147	monocytogenes dissemination in the host. (Demonstrated in mice) CLEC7A (Dectin-1) is a C-type lectin receptor that
18428	CLEC7A	15956283	binds beta-glucan and is the primary receptor on macrophages for phagocytosis of various fungi.
18428	CLEC7A	17698636	CLEC7A synergizes with TLR2 to increase pro- inflammatory cytokine responses to pathogens. CLEC7A has a critical role in the innate immune
18428	CLEC7A	19633936	response against Mycobacterium tuberculosis in non- phagocytic cells in type II airway epithelial cells. CLEC7A-dependent immune-cell recognition of beta glucan on the fungal cell-wall is modulated by the Candida albicans ERK-like-1 (CEK1)-mediated MAPK pathway, disruption of which causes enhanced cell-wall beta-glucan exposure triggering immune responses
18428	CLEC7A	20100861	more efficiently than wild-type yeast.

CLEC7A, a C-type lectin receptor, is a pattern recognition receptor critical for immune responses to fungi. CLEC7A is coupled to SYK kinase and signals via CARD9 to activate NFKB, which in turns induces both innate and adaptive immunity.

18428 CLEC7A 21267996
 CLEC7A is expressed mainly in dendritic cells and macrophages. CLEC7A recognizes beta-glucans with its carbohydrate recognition domain and transduces signals through its ITAM-like motif in the cytoplasmic region, which recruits SYK and initiates the CARD9/NFkB signalling cascade. CLEC7A acts as an extracellular sensor for fungi and

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CLEC7A

CLEC7A

Hamp

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HAMP

HAMP

HAMP

21572427

21685415

22544439

21572427

22267217 CLEC/A acts as an extracellular sensor for fungi and mycobacteria that induce both IL1B production and maturation for protective immunity.

22543832 The functional activity of CLEC7A in mucosal immunity to Candida albicans is dependent on the genetic background of the host. (Demonstrated in mice) Hamp is a peptide hormone that regulates iron homeostasis and acts as an antimicrobial peptide, functioning in modulating acute inflammatory 20530874 responses by mediating transcriptional changes.

> Hamp (hepcidin), an iron regulatory hormone, inhibits the liver-stage infection in malarial parasitemia by arresting Plasmodium sporozoites in liver hepatocytes and prevents their development into blood-stage parasites.

Hamp expression in LPS or Tnfa-stimulated peripheral blood leukocytes is dependent on the NFkB signalling pathway. (Demonstrated in human)

Hamp expression is induced by Tlr2/Tlr4 ligands and may regulate pro-inflammatory cytokine production through maintenance of iron homeostasis in macrophages.

HAMP (hepcidin), an iron regulatory hormone and antimicrobial peptide, inhibits the liver-stage infection in malarial parasitemia by arresting Plasmodium sporozoites in liver hepatocytes and prevents their development into blood-stage parasites. (Demonstrated in murine model)

HAMP expression in LPS or TNFA-stimulated peripheral blood leukocytes is dependent on the NFkB signalling pathway.

HAMP expression is induced by TLR2/TLR4 ligands and may regulate pro-inflammatory cytokine production through maintenance of iron homeostasis in macrophages. (Demonstrated in mice)

214493Rhbdf222550345Rhbdf2 is required for the secretion of Tnf in<br/>macrophages.

69739	RHBDF2	22550345	RHBDF2 is required for the secretion of TNF in macrophages. (Demonstrated in mice) Plscr1 regulates Tlr9 trafficking to endosomal compartment and plays an important role in the
191069	Plscr1	22453241	induction of type I IFN response in plasmocytoid dendritic cells. PLSCR1 regulates TLR9 trafficking to endosomal compartment and plays an important role in the
60169	PLSCR1	22453241	induction of type I IFN response in plasmocytoid dendritic cells. DUSP1 inhibits MAPK pathways and decreases TLR
57812	DUSP1	18504304	signalling. DUSP1 is an essential feedback regulator of the innate immune response, it plays a critical role in preventing septic shock and multi-organ dysfunction during pathogenic infection and plays a pivotal role in the
57812	DUSP1	19436832	deactivation of MAPK1 and MAPK8. DUSP1 is a negative regulator of MAPK-dependent
57812	DUSP1	21959016	induction of IL6 and IL8 in response to the coronavirus infectious bronchitis virus (IBV). DUSP1 antagonizes p38 MAPK activity to induce IL12B expression, and may play a role in the
57812	DUSP1	22464096	development of Th1 type immune response and anti- microbial defence. (Demonstrated in mice) Tlr4::Ly96 functions as intracellular LPS sensor and
135441	Ly96	21712422	triggers a unique set of LPS responses upon recognition of phagocytosed bacteria in macrophages. Amino acid Phe126 of Ly96 acts as a hydrophobic switch driving agonist-dependent contacts necessary for Tlr4 dimerization and activation. (Demonstrated in
135441	Ly96	22433852	human) Morphine binds to Ly96, triggering Tlr4
135441	Ly96	22474354	oligomerization and induces neuroinflammation within the central nervous system. (Demonstrated in human) LY96 (MD-2) recognizes only exogenous pathogen-
25842	LY96	19840871	associated molecular patterns (PAMPs) when complexed with TLR2-CD14 or TLR4-CD14. LY86 and CD180 interact directly with the TLR4
25842	LY96	15852007	signalling complex, inhibiting its ability to bind microbial ligand. LY96 is physically associated with TLR4 on the cell
25842	LY96	10359581	surface and confers responsiveness to lipopolysaccharide (LPS). TLR4::LY96 functions as intracellular LPS sensor and triggers a unique set of LPS responses upon
25842	LY96	21712422	recognition of phagocytosed bacteria in macrophages. (Demonstrated in murine model) Amino acid Phe126 of LY96 acts as a hydrophobic
25842	LY96	22433852	switch driving agonist-dependent contacts necessary for TLR4 dimerization and activation.

			Morphine binds to LY96, triggering TLR4 oligomerization and induces neuroinflammation within
25842	LY96	22474354	the central nervous system. LILRA2-activation of monocytes is distinct from LPS activation, as assessed by the secreted cytokine profile.
409454	LILRA2	22479404	LILRA2 cross-linking also results in inhibition of FCGR1A-dependent phagocytosis in monocytes. Tyrosine phosphorylation of the E3 ubiquitin ligase
202909	Trim21	22479513	Trim21 positively regulates interaction with Irf3 and hence Trim21 activity. Clec9a is restricted to dendritic cells, and specifically recognizes filamentous actin from necrotic cells. Clec9a ligand engagement is necessary for the priming
191954	Clec9a	22483800	of cytotoxic T cells against necrotic cell antigens. (Demonstrated in human) CLEC9A is restricted to dendritic cells, and specifically recognizes filamentous actin from necrotic cells. CLEC9A ligand engagement is necessary for the
18377	CLEC9A	22483800	priming of cytotoxic T cells against necrotic cell antigens.
157012	Pacsin1	22488361	Pacsin1 regulates the Tlr7/9-mediated type I interferon response in plasmacytoid dendritic cells.
83573	PACSIN1	22488361	PACSIN1 regulates the TLR7/9-mediated type I interferon response in plasmacytoid dendritic cells. Irak3 dampens the innate immune response in airway
192499	Irak3	22154382	epithelial cells by inhibiting Tlr2 signalling in an Il13- dependent manner. (Demonstrated in human)
192499	Irak3	22492852	Irak3 impairs host defence during pneumococcal pneumonia at the primary site of infection. IRAK3 upregulation by lipopolysaccharide (LPS) is largely dependent on TNF-alpha and LPS tolerance in
45584	IRAK3	17982103	human endotoxemia models is associated with IRAK3 up-regulation. IRAK3 selectively attenuates Pam3CSK4-induced
45584	IRAK3	17379480	MAPK14 (p38) activation through an IRAK1 independent and MKP1 dependent pathway. IRAK3 is part of the interleukin-1 receptor-associated kinase (IRAK) family of proteins that mediate
45584	IRAK3	12620219	activation of nuclear factor-kappaB (NF-kappaB) and mitogen-activated protein kinase (MAPK) pathways. IRAK3 is one of the negative regulators that contribute to the attenuation of NF-kappaB activation and it
45584	IRAK3	19809574	negatively regulates the alternative NF-kappaB pathway in a ligand-specific manner. IRAK3 induction and inhibition of kinase activity of
45584	IRAK3	14660668	IRAK1 are crucial to peptidoglycan-induced tolerance in macrophages. IRAK3 regulates Toll-like receptor (TLR) signalling and innate immune homeostasis where IRAK3 is
45584	IRAK3	12150927	induced upon TLR stimulation and negatively regulates TLR signalling.

45584	IRAK3	20042589	IRAK3 is critical to preventing deleterious neutrophil- dependent lung injury during influenza infection of the respiratory tract.
15564	interest	20042307	IRAK3 is strongly expressed in resting alveolar macrophages and blunts TNF expression to protect the host from the injurious effect of bacterial-induced inflammation. The inhibitory function of IRAK3 is abolished when it is cleaved by CASP6, which allows the induction of the NFKB pathway and the expression
45584	IRAK3	21098228	of TNF. IRAK3 dampens the innate immune response in airway
45584	IRAK3	22154382	epithelial cells by inhibiting TLR2 signalling in an IL13-dependent manner. IRAK3 impairs host defence during pneumococcal
45584	IRAK3	22492852	pneumonia at the primary site of infection. (Demonstrated in mice) Eps8 is a key regulator of the LPS-induced Tlr4-
195386	Eps8	22493489	Myd88 interaction and directly contributes to phagocytosis in macrophages. EPS8 is a key regulator of the LPS-induced TLR4-
21563	EPS8	22493489	MYD88 interaction and directly contributes to phagocytosis in macrophages. (Demonstrated in mice)
163574	Atf4	22496230	Murine cytomegalovirus targets transcription factor ATF4 to exploit the unfolded protein response Murine cytomegalovirus targets transcription factor
8409	ATF4	22496230	ATF4 to exploit the unfolded protein response (Demonstrated in mice) Nfat5 is a transcription factor that is upregulated by hypertonicity. Activated Nfat5 enhances LPS-mediated NFkB activity by complexing with NFkB and kB
190689	Nfat5	20685965	elements in downstream genes. Nfat5 expression is strongly induced by Mycobacterium tuberculosis, and plays an essential role in stimulating HIV replication in co-infected
190689	Nfat5	22496647	macrophages. (Demonstrated in human) NFAT5 is a transcription factor that is upregulated by hypertonicity. Activated NFAT5 enhances LPS- mediated NFkB activity by complexing with NFkB and kB elements in downstream genes. (Demonstrated
39265	NFAT5	20685965	in murine model) NFAT5 expression is strongly induced by Mycobacterium tuberculosis, and plays an essential
39265	NFAT5	22496647	role in stimulating HIV replication in co-infected macrophages. Ehmt2 catalyzes di-methylation of histone H3 at the promoter of interferon (IFN) and IFN-inducible antiviral genes as an epigenetic silencing mechanism in fibroblasts. Ablation of Ehmt2 results in phenotypic conversion to potent IFN-producing cells and renders
174783	Ehmt2	22412156	resistance to RNA viruses.
			EHMT2 catalyzes di-methylation of histone H3 at the promoter of interferon (IFN) and IFN-inducible antiviral genes as an epigenetic silencing mechanism in fibroblasts. Ablation of EHMT2 results in phenotypic
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79489	EHMT2	22412156	conversion to potent IFN-producing cells and renders resistance to RNA viruses. (Demonstrated in mice) TSC1 inhibits TLR response and endotoxin tolerance
90470	TSC1	22412198	through repression of mTORC1 and JNK1/2 signalling pathways. (Demonstrated in mice) IRF5 is activated by both TBK1 and MYD88 to form a
40393	IRF5	18332133	homodimer that binds to and activates transcription of type I interferon and inflammatory cytokine genes. IRF5 can act as both an activator and a repressor of interferon (IFN) gene induction dependent on the IRF- interacting partner, and may be a part of the regulatory network that ensures timely expression of the
40393	IRF5	12600985	immediate early inflammatory genes. Phosphorylation of IRF5 at ser451 and ser462 is the
40393	IRF5	22412986	primary trigger of IRF5 function in nuclear accumulation, transcription and apoptosis. Sftpa1 is a carbohydrate pattern recognition molecule of innate immunity, that significantly enhances phagocytosis and killing of Aspergillus fumigatus, a
151763	Sftpal	20413160	pathogenic fungus, by neutrophils and macrophages. Sftpa1 binds to Mycobacterium avium lipid and it promotes the acclutination of the pathogen. The
151763	Sftpa1	21821801	promotes the agglutination of the pathogen. The presence of Sftpa1 inhibits the growth of M. avium in culture. (Demonstrated in human) Sftpa1 confers protection in lung epithelium against the
151763	Sftpa1	22418431	cytotoxic effects of human beta-defensin 3, DEFB103B. Rubicon is a physiological feedback inhibitor of pattern
154810	1700021K19Ril	22423967	recognition receptor signalling, preventing unbalanced proinflammatory responses. (Demonstrated in human) Rubicon positively regulates the NADPH oxidase complex to induce phagosomal trafficking of p22phox- gp91phox (which are subunits of NADPH oxidase),
154810	1700021K19Ril	22423966	and the induction of reactive oxygen species and inflammatory cytokines. (Demonstrated in human) RUBICON is a physiological feedback inhibitor of
71731	KIAA0226	22423967	pattern recognition receptor signalling, preventing unbalanced proinflammatory responses. RUBICON positively regulates the NADPH oxidase complex to induce phagosomal trafficking of p22phox- gp91phox (which are subunits of NADPH oxidase), and the induction of reactive oxygen species and
71731	KIAA0226	22423966	inflammatory cytokines. ZBP1 is a cytosolic DNA sensor that upon binding to DNA, associates with IRF3 transcription factor and
82385	ZBP1	17618271	TBK1 serine/threonine kinase, selectively enhancing DNA mediated induction of type-I-IFN.

			ZBP1 is a cytosolic pattern recognition receptor of double-stranded DNA and is essential for IRF3 activation and interferon beta expression triggered by human cytomegalovirus (HCMV), as well as being sufficient to enhance HCMV-stimulated beta interferon
82385	ZBP1	19846511	transcription and secretion. ZBP1 (DAI) enhanced HIV-1 replication, which is largely impaired by mutations at kappaB sites in HIV-1 long terminal repeat (LTR) or by suppressing
82385	ZBP1	20599623	activation of NF-kappaB. ZBP1 interacts with RIPK3 to mediate virus-induced
82385	ZBP1	22423968	necrosis. (Demonstrated in mice) RIPK3 forms a complex with TICAM1 upon Toll-like receptors (TLR) 3 and 4 activation resulting in RIPK3- dependent but TNF-independent necrosis in
3902	RIPK3	22123964	macrophages. (Demonstrated in mouse) RIPK3 interacts with ZBP1 to mediate virus-induced
3902	RIPK3	22423968	necrosis. (Demonstrated in mice)
145299	Tmed7	22426228	Tmed7 inhibits LPS-mediated Tlr4 signalling in late endosomes. (Demonstrated in human) TMED7;TICAM2 (TICAM2) splice variant TAG
37819	'MED7;TICAM	19412184	negatively regulates the adapter MYD88-independent TLR4 pathway by displacing TICAM1 adapter from TICAM2. TICAM2 and TICAM1 both function in LPS-TLR4 signaling to regulate the MYD88-independent pathway
37819	'MED7;TICAM	14517278	during the innate immune response to lipopolysaccharide (LPS). TICAM2 and TICAM1 form an adapter complex that
37819	MED7;TICAM	14519765	plays a crucial role for LPS-TLR4-mediated activation of IFN-beta. TMED7;TICAM2 is an adapter protein for both TLR4
37819	MED7;TICAM	17277119	and TLR2/6 signaling in synovial fibroblasts, HUVECs, and murine embryonic fibroblasts.
37819	MED7;TICAM	14556004	TICAM2 provides specificity for the MYD88- independent component of TLR4 signalling.
37819	MED7;TICAM	22426228	TMED7 inhibits LPS-mediated TLR4 signalling in late endosomes. Illrap is recruited to the Illb::Illr1 ligand-receptor
150439	Illrap	22426547	complex to initiate the II1b signalling cascade. (Demonstrated in human) IL1RAP is recruited to the IL1B::IL1R1 ligand-
69682	IL1RAP	22426547	receptor complex to initiate the IL1B signalling cascade. IL1R1 is a substrate for presenilin-dependent gamma-
64088	IL1R1	18996842	secretase cleavage and intramembrane proteolysis may be a control mechanism for IL1R1-mediated signalling. IL1R1 is essential for TLR9-dependent activation of tumour necrosis factor receptor-associated factor 3
64088	IL1R1	21115691	(TRAF3) and for production of the anti-inflammatory cytokines IL-10 and type I interferon (IFN).

64088	IL1R1	22426547	IL1R1 is the primary receptor for the inflammatory cytokine IL1B.
212144	Sdc4	22427536	Sdc4 acts in the early inflammatory response to LPS and functions to limit the extent of pulmonary inflammation and lung injury.
77503	SDC4	22427536	SDC4 acts in the early inflammatory response to LPS and functions to limit the extent of pulmonary inflammation and lung injury. (Demonstrated in mice)
			Tlr7 and Tlr8 are translocated from the endoplasmic reticulum to the endosome in the presence of antiphospholipid antibodies, as a consequence, plasmacytoid dendritic cells become dramatically sensitized to Tlr7/8 agonists and this may play a role in
185523	Tlr8	21734241	systemic autoimmunity. Tlr8 binding of HIV ssRNA induces endosomal acidification and chromatin remodeling at the TNF-
185523	Tlr8	22393042	alpha promoter to promote TNF-alpha release in infected macrophages. (Demonstrated in human) Hmgb1 is an endogenous Tlr4 ligand in macrophages
209311	Hmgb1	21372296	and its release in wounds initiates Tlr4-dependent responses that contribute to neovascularization. Hmgb1 plays a key regulatory role in polymorphonuclear neutrophil (PMN) recruitment to
209311	Hmgb1	21860212	inflammatory tissues. Low concentrations of Hmgb1 (50-100 ng/ml) reduce baseline PMN migration as well as formyl-methionyl-leucyl-phenylalanine- and II8-induced PMN chemotaxis, whereas higher Hmgb1 concentrations (5000 ng/ml) have a chemoattractant effect on PMN through II8 production . (Demonstrated in human)
			Hmgb1 has a pathogenic role in arthritis, where in complex with lipopolysaccharide, Il1a or Il1b, Hmgb1 boosts the production of proinflammatory cytokines and Mmp3 as demonstrated in synovial fibroblasts
209311	Hmgb1	21871094	from rheumatoid arthritis and osteoarthritis patients. (Demonstrated in human) Nuclear Hmgb1 translocates to the cytoplasm in LPS-
209311	Hmgb1	22396017	stimulated macrophages to potentiate inflammatory responses. (Demonstrated in mice) Cd209a has a role in the regulation of inflammation in
130586	Cd209a	20130211	a model of experimental colitis and is a critical innate factor in response to LPS. Cd209a is involved in cell-cell contact signalling between activated apoptotic lymphocytes and dendritic
130586	Cd209a	22396536	cells (DC) during the maturation of DCs. (Demonstrated in human) CD209, upon pathogen binding, induces an intracellular signalling pathway with a central role for the serine/threonine kinase RAF1 and modulates TLR-
23414	CD209	18998127	induced activation.

23414	CD209	22396536	CD209 is involved in cell-cell contact signalling between activated apoptotic lymphocytes and dendritic cells (DC) during the maturation of DCs. Samhd11 is specifically targeted for degradation by
211114	Samhd1	21720370	human immunodeficiency virus 1 (HIV-1) viral protein VPX, and mutations in SAMHD11 causes Aicardi- Goutieres syndrome, which mimics symptoms of congenital viral infection. (Demonstrated in human) Samhd1 is expressed in dendritic and myeloid cells and acts as an anti-retroviral protein that inhibits the early stages of the viral life cycle. Silencing of SAMHD1
211114	Samhd1	21613998	leads to significant accumulation of human immunodeficiency virus 1 (HIV-1) DNA in infected cells. (Demonstrated in human) Samhd1 inhibits HIV replication by depleting intracellular dNTPs, which are required for viral
211114	Samhd1	22327569	reverse transcriptase to synthesize viral DNA. (Demonstrated in human) Ntn1 inhibits Ccl2 and Ccl19-mediated chemotaxis in
188640	Ntn 1	22231519	macrophages and promotes atherosclerosis by retaining macrophages in the artery wall. NTN1 inhibits CCL2 and CCL19-mediated chemotaxis in macrophages and promotes atherosclerosis by
29663	NTN1	22231519	retaining macrophages in the artery wall. (Demonstrated in mice)
134049	Nampt	22377803	Nampt secretion is enhanced by extracellular ATP in LPS-primed monocytes. (Demonstrated in human) NAMPT secretion is enhanced by extracellular ATP in
35259	NAMPT	22377803	LPS-primed monocytes.
80463	AGER	22386596	AGER is a native receptor for complement component C1QA.
166648	Jak3	22359619	Jak3 inhibition is sufficient to counteract lung injury and protect immunocytes from hypercytokinemia following challenge with avian influenza H5N1 hemagglutinin. JAK3 is a primary response gene for IL6 in macrophage differentiation and ectopic overexpression of JAK3 accelerates monocytic differentiation of
37201	JAK3	14976041	normal mouse bone marrow cells stimulated with cytokines. JAK3 inhibition is sufficient to counteract lung injury
37201	JAK3	22359619	and protect immunocytes from hypercytokinemia following challenge with avian influenza H5N1 hemagglutinin. Sharpin is an essential adaptor downstream of Ikbkg in the TLR signalling pathway. A mutation in the Sharpin
154756	Sharpin	21709223	gene impairs II12 production in TLR-stimulated macrophages. Sharpin deficient dendritic cells show reduced
154756	Sharpin	22348129	inflammatory cytokine production, defective NF-kB signalling and skewing towards a Th2-response.

			SHARPIN is an essential adaptor downstream of
			IKBKG in the TLR signalling pathway. A mutation in
			the SHARPIN gene impairs IL12 production in TLR-
40065	SHARPIN	21709223	stimulated macrophages.
			SHARPIN deficient dendritic cells show reduced
			inflammatory cytokine production, defective NF-kB signalling and skewing towards a Th2-response.
40065	SHARPIN	22348129	(Demonstrated in mice)
			Camkk2-null macrophages exhibited deficiency to
			spread, phagocytize bacteria and synthesize cytokines
198659	Camkk2	22334678	in response to the Toll Like Receptor 4 (TLR4) agonist lipopolysaccharide (LPS).
198039	Callikk2	22334078	CAMKK2-null macrophages exhibited deficiency to
			spread, phagocytize bacteria and synthesize cytokines
			in response to the Toll Like Receptor 4 (TLR4) agonist
61470	CAMKK2	22334678	lipopolysaccharide (LPS). (Demonstrated in mice)
			WNT3A in human chondrocytes counteracts IL1B induced NF-kB mediated matrix metalloproteinases
107120	WNT3A	22328140	expression in a negative feedback loop.
			Was interacts with Btk to induce the LPS signalling
130090	Was	22253930	cascade in macrophages.
(2702	WA C	22252020	WAS interacts with BTK to induce the LPS signalling
63703	WAS	22253930	cascade in macrophages. (Demonstrated in mice) MIR16-2 expression is induced upon LPS stimulation
			and acts to promote NF-kB-mediated transcription of
126583	MIR16-2	22292036	IL8 by suppressing the translation of SMRT.
			MIR16-1 expression is induced upon LPS stimulation
127053	MIR16-1	22292036	and acts to promote NF-kB-mediated transcription of
12/035	MIK10-1	22292030	IL8 by suppressing the translation of SMRT. PYCARD protein contains CARD and Pyrin domains
28012	PYCARD	18487086	and is required for assembly of inflammasome.
			PYCARD prevents oligomerization of CASP1
			mediated by RIPK2 by out-competing RIPK2 for
			binding, and thus, preventing CASP1 autoactivation. PYCARD also recruits CASP1 into PYCARD-formed
28012	PYCARD	14634131	cytosolic specks, separating it from RIPK2.
	-		PYCARD directs CASP1 away from RIPK2-mediated
			NF-kappaB activation, toward CASP1-mediated
28012	PYCARD	16585594	processing of pro-IL1B by interfering with CASP1- RIPK2 interaction.
28012	PICARD	10383394	PYCARD binds to AIM2 to form a CASP1 and NF-
28012	PYCARD	19158675	kappaB activating inflammasome.
			PYCARD (ASC) splice variant protein (vASC) lacking
00010		10550050	the PGR domain regulates IL1B release and aggregates
28012	PYCARD	19759850	differently from intact PYCARD. PYCARD is a component of the inflammasome and is
			required for inflammation in acute pancreatitis.
28012	PYCARD	21439959	(Demonstrated in murine model)
			NLRP3/PYCARD inflammasome activation following
			human respiratory syncytial virus infection is
28012	PYCARD	22295065	dependent on the activation of TLR2/MYD88/NF-kB and reactive oxygen species/potassium efflux.
20012	TICAND	22275005	and reactive oxygen species/polassium emux.

			Frem1 selectively amplifies NF-kB responses, such as cell survival and inflammatory responses, through
159685	Frem1	22262840	changes in receptor conformation and adapter protein recruitment. (Demonstrated in human) FREM1 selectively amplifies NF-kB responses, such as cell survival and inflammatory responses, through
50859	FREM1	22262840	changes in receptor conformation and adapter protein recruitment. Usp4 is a potent negative regulator of Tlr/Il1r-
199676	Usp4	22262844	signalling through deubiquitination of Traf6 to prevent the activation of inflammatory responses. USP4 is a potent negative regulator of TLR/IL1Rsignalling through deubiquitination of
34595	USP4	22262844	TRAF6 to prevent activation of inflammatory responses.
38275	DCD	19014393	Activates normal human keratinocytes and display in vitro microbicidal activities against bacteria and viruses DCD encodes the anionic amphiphilic peptide DCD-1L that interacts with negatively charged bacterial
38275	DCD	22262861	phospholipids to form ion channels in the bacterial membrane. Following TNF stimulation, PRKCD phosphorylates
40082	PRKCD	19150425	TRAF2 leading to CHUK (IKK alpha) and IKBKB (IKK beta) recruitment to the TNF receptor PRKCD is an essential signalling kinase in C-type
40082	PRKCD	22265677	lectin receptor-mediated innate immunity and host protection. (Demonstrated in mice) Fstl1 acts as an endogenous Tlr4 agonist. Similar to
159696	Fstl1	22265692	LPS, Fstl1 induces tolerance to subsequent LPS stimulations. FSTL1 acts as an endogenous TLR4 agonist and has the ability to induce IL6 and IL8 production. Similar to
51777	FSTL1	22265692	LPS, FSTL1 induces tolerance to subsequent LPS stimulations. The transcription factor RORA is critical for the development of nuocytes and the mounting of innate
14920	RORA	22267218	<ul><li>type 2 immunity against parasitic worms.</li><li>(Demonstrated in mice)</li><li>DEFA1 (HNP-1) is a cationic alpha-defensin peptide</li></ul>
6083	DEFA1	19024344	present in human neutrophils and alpha-defensins have multiple functions in the immune system. Defensins, such as DEFA1, contribute to innate immunity through diverse actions, including microbial
6083	DEFA1	19897717	killing and high concentrations are present in the lung in response to inflammation. DEFA1, an alpha-defensin, binds to the cell wall precursor lipid II and reduction of lipid II levels in the bacterial membrane significantly reduces bacterial killing, suggesting that the inhibition of cell wall synthesis is a novel antibacterial mechanism of this
6083	DEFA1	20214904	important class of host defence peptides.

			The dimerization of DEFA1 is crucial for the ability of
6083	DEFA1	22270360	the alpha defensin to kill S. aureus, inhibit anthrax lethal factor and bind HIV-1 protein gp120. DEFA1 activates platelets and induces the formation of
6083	DEFA1	22268819	amyloid-like proteins, which entrap bacteria and fungi. Trib2 expression is induced by Tlr5 stimulation to
129207	Trib2	22271508	inhibit NF-kB signalling in epithelial cells. TRIB2 expression is induced by TLR5 stimulation to
30297	TRIB2	22271508	inhibit NF-kB signalling in epithelial cells. Apoal has anti-inflammatory properties in
			macrophages through the stabilization of ATP-binding cassette transporter A1 (Abca1) and the down- regulation of Tlr4 signalling pathway. (Demonstrated
161189	Apoa1	22271762	in human) APOA1 is a serum apolipoprotein that induces antiatherogenic efflux of macrophage cholesterol and is anti-inflammatory due to its ability to neutralize bacterial lipopolysaccharide (LPS). APOA1 also signals in the macrophage through Toll-like receptor (TLR)2, TLR4, and CD14, utilizing MYD88- dependent and -independent pathways, to activate
72644	APOA1	20519121	nuclear factor-kappaB and induce cytokines. APOA1 has anti-inflammatory properties in macrophages through the stabilization of ATP-binding cassette transporter A1 (ABCA1) and the down-
72644	APOA1	22271762	regulation of TLR4 signalling pathway. Jak2 inhibition prevents innate immune responses and rescues animals from sepsis by specifically preventing LPS-induced STAT3 tyrosine phosphorylation without affecting serine phosphorylation in macrophages and by preventing the activation of the canonical p65RelA/p50NF-kappaB1 pathway but not the other
156977	Jak2	20393690	NF-kappaB proteins. Jak2 functions downstream of Tlr2-stimulation to
156977	Jak2	22218715	induce the expression of the monocyte chemoattractant protein 1, Ccl2. JAK2 binds to the majority of the known members of the cytokine family of receptors and ligand-receptor binding leads to activation of the associated JAK2 molecules, resulting in rapid autophosphorylation of multiple tyrosines within JAK2. Tyrosine 813 is a site of autophosphorylation in JAK2 and is the SH2-B beta- binding site within JAK2 that is required for SH2 P
47039	JAK2	15121872	binding site within JAK2 that is required for SH2-B beta to enhance activation of JAK2. JAK2 and JAK1 tyrosine kinases physically associate with the gamma chain and beta chain of the interleukin 2 (IL-2) receptor , respectively, suggesting that regulation of the kinases may be linked to IL-2-induced
47039	JAK2	8041779	signal transduction.

JAK2 inhibition prevents innate immune responses and rescues animals from sepsis by specifically preventing LPS-induced STAT3 tyrosine phosphorylation without affecting serine phosphorylation in macrophages and by preventing the activation of the canonical p65RelA/p50NF-kappaB1 pathway but not the other NF-kappaB proteins.

JAK2 functions downstream of TLR2-stimulation to induce the expression of the monocyte chemoattractant protein 1, CCL2. (Demonstrated in mice)

Hsp90b1 on the surface of polymorphonuclear leukocytes (PMNs) serves as a receptor for Escherichia coli K1 (EC-K1) entry. EC-K1 exploits surface-expressed Hsp90b1 in PMNs to prevent oxidative burst for the onset of neonatal meningitis.

Hsp90b1is an essential endoplasmic reticulum182078Hsp90b122223641chaperone protein for Toll-like receptors and integrins.

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HSP90B1HSP90B1 amplifies innate and adaptive immuneHSP90B116754684responses via interaction with TLR2 and TLR4 ligands.<br/>Absence of HSP90B1 results in a defect in the<br/>formation of some cell surface receptors including<br/>Toll-like receptors (TLRs) which, as a result, are<br/>retained intracellularly.

HSP90B1 is an endoplasmic reticulum master chaperone for multiple TLRs and is also essential for expression of multiple hematopoietic system-specific integrins.

HSP90B1, an endoplasmic reticulum chaperone, is required in vesicular stomatitis virus (VSV) infection and in innate immunity via Toll-like receptors (TLRs). TLR-mediated immunity maintains the broad host range of the envelope glycoprotein of VSV (VSV-G) by positively selecting for the ubiquitous expression of HSP90B1.

HSP90B1 chaperones multiple Toll-like receptors (TLRs), but not TLR3, in a manner that is dependent on another endoplasmic reticulum (ER) luminal protein, CNPY3.

HSP90B1 on the surface of polymorphonuclear leukocytes (PMNs) serves as a receptor for Escherichia coli K1 (EC-K1) entry into the cell. EC-K1 exploits surface-expressed HSP90B1 in PMNs to prevent oxidative burst for the onset of neonatal meningitis. (Demonstrated in mice)

HSP90B1 is an essential endoplasmic reticulum chaperone protein for Toll-like receptors (TLR) and integrins. (Demonstrated in mice)

Trpm2Trpm2Trpm2Trpm2Trpm2167747Trpm221709234inefficient innate immune response.

167747	Trpm2	22101731	Trpm2 inhibits reactive oxygen species (ROS) production in phagocytic cells and prevents endotoxin-induced lung inflammation in mice.
10//4/	1101112	22101751	Trpm2 deficient mice are extremely susceptible to infection with Listeria monocytogenes, exhibiting an inefficient innate immune response. (Demonstrated in
5302	TRPM2	21709234	murine model) TRPM2 inhibits reactive oxygen species (ROS)
5302	TRPM2	22101731	production in phagocytic cells and prevents endotoxin- induced lung inflammation. (Demonstrated in mice) Cyld is a deubiquitinase that act as a negative regulator
179176	Cyld	21498625	of TLR3 induction in response to LPS. (Demonstrated in human) Cyld plays a key role in Type I IFN receptor signalling during vesicular stomatitis virus (VSV) infection. In the channes of Cyld. USN here is ineffective in the
179176	Cyld	21946435	the absence of Cyld, IFN-beta is ineffective in the induction of antiviral genes. The E3 ligase Itch and deubiquitinase Cyld act together
179176	Cyld	22057290	to regulate Tak1 and inflammation.
210517	Itch	22057290	The E3 ligase Itch and deubiquitinase Cyld act together to regulate Tak1 and inflammation. ITCH is an E3 ligase that negatively regulates
68143	ITCH	18246070	inflammatory signalling pathways by controlling the function of the ubiquitin-editing enzyme TNFAIP3. ITCH is a HECT domain-containing E3 ligase that negatively regulates MAVS-mediated antiviral
68143	ITCH	19881509	response by catalyzing the K48-linked polyubiquitination and degradation of MAVS. The E3 ligase ITCH and deubiquitinase CYLD act
68143	ITCH	22057290	together to regulate TAK1 and inflammation. IL17C is an essential autocrine cytokine that regulates
46304	IL17C	21993848	innate epithelial immune responses. Il17re is a receptor specific to Il17c that regulates early
176930	Il17re	21993849	innate immunity to intestinal pathogens in colon epithelial cells. IL17RE is a receptor specific to IL17C that regulates
16793	IL17RE	21993849	early innate immunity to intestinal pathogens in colon epithelial cells. (Demonstrated in mice) IL9 production is largely restricted to innate lymphoid cells during papain-induce lung inflammation and IL9
45710	IL9	21983833	serves as an important bridging link to induce type 2 helper T cell responses. (Demonstrated in mice) Lrrk2 acts as a potent negative regulator of the transcription factor NFAT, and plays an important role in modulating inflammatory bowel disease. Lrrk2
175625	Lrrk2	21983832	deficiency conferred enhanced susceptibility to experimental colitis in mice. LRRK2 acts as a potent negative regulator of the
27257	LRRK2	21983832	transcription factor NFAT, and plays an important role in modulating inflammatory bowel disease.

			Class III phosphatidylinositol 3-kinases (PI3K) are required for downstream ARF6 regulation of CpG
2711	PIK3C3	22170068	oligodeoxynucleotide uptake and thus have a role in TLR9-mediated immune signalling. Arf6 has a pivotal role in Tlr9-mediated immune
144539	Arf6	22170068	signaling by regulating the cellular uptake of CpG oligodeoxynucleotides. IL13 is a pro-M2/Th2 cytokine that induces alternative
42599	IL13	21097505	activation of macrophages. IL13 associates with IL13RA1 to activate the transcription factor STAT6. IL13 dampens the innate immune response in airway
42599	IL13	22154382	epithelial cells via IRAK3-mediated inhibition of TLR2 signalling.
191235	Нр	22156194	Haptoglobin activates innate immunity to enhance acute transplant rejection.
41488	HP	22156194	Haptoglobin activates innate immunity to enhance acute transplant rejection. (Demonstrated in mouse) Unc5cl is a factor in epithelial inflammation where it
190027	Unc5cl	22158417	activates the pro-inflammatory IRAK signalling cascade in a Myd88-independent manner. UNC5CL is a factor in epithelial inflammation where it
86449	UNC5CL	22158417	activates the pro-inflammatory IRAK signalling cascade in a MYD88-independent manner. Plcg2, 1,4,5-triphosphate and intracellular calcium are required for the LPS-induced innate immune response
195070	Plcg2	22158869	pathway, where release of intracellular calcium mediates Tlr4 trafficking and subsequent activation of Irf3. PLCG2, 1,4,5-triphosphate and intracellular calcium are required for the LPS-induced innate immune
43723	PLCG2	22158869	response pathway, where release of intracellular calcium mediates TLR4 trafficking and subsequent activation of IRF3. (Demonstrated in mouse) Cnot4 enhances JAK/STAT pathway-dependent gene expression by positively regulating IFN-gamma- and
135345	Cnot4	22159038	IL4-induced STAT-mediated gene responses. (Demonstrated in human) CNOT4 enhances JAK/STAT pathway-dependent gene
42604	CNOT4	22159038	expression by positively regulating IFN-gamma- and IL4-induced STAT-mediated gene responses. Unc93b1 is a resident endoplasmic reticulum protein that interacts with Tlr11 to regulate the dendritic cell
128581	Unc93b1	21097503	<ul><li>activation in response to T. gondii profilin and parasitic infection.</li><li>Unc93b1 controls homeostatic Tlr7 activation by balancing Tlr9 to Tlr7 trafficking from endoplasmic reticulum to endolysosomes. D34A mutation in</li></ul>
128581	Unc93b1	21683627	Unc93b1 causes Tlr7-dependent systemic lethal inflammation.

			Unc93B1 is a multitransmembrane endoplasmic
			reticulum (ER)-resident protein that controls
			homeostatic Tlr7 activation by balancing Tlr9 to Tlr7 trafficking. Mice harboring a D34A mutation in
			Unc93b1 show Tlr7-dependent, systemic lethal
			inflammation.
128581	Unc93b1	21683627	innanniation.
120501	0110/301	21005027	Unc93b1 participates in intracellular trafficking and
			signalling for all nucleotide-sensing Toll-like receptors
128581	Unc93b1	22164301	(TLRs). (Demonstrated in human)
			UNC93B1 is a polytopic membrane protein that
			delivers the nucleotide sensing receptors TLR7 and
			TLR9 from the endoplasmic reticulum to
61027	UNC93B1	18305481	endolysosomes.
			UNC93B1 biases Toll-like receptor responses to
			nucleic acid in dendritic cells toward DNA- but against
61027	UNC93B1	19451267	RNA-sensing.
			UNC93B1 associates with Toll-Like Receptor (TLR) 3,
			TLR7 and TLR9, mediating their translocation from
			the endoplasmic reticulum to the endolysosome, hence
			allowing proper activation by nucleic acid ligands.
			UNC93B1 has a critical role on induction of IL-
			12/IFN-gamma production as well as autonomous
61027	UNC93B1	20865117	control of Toxoplasma replication by macrophages.
			UNC93B1 controls homeostatic TLR7 activation by
			balancing TLR9 to TLR7 trafficking from endoplasmic
			reticulum to endolysosomes. D34A mutation in
(1005			UNC93B1 causes TLR7-dependent systemic lethal
61027	UNC93B1	21683627	inflammation. (Demonstrated in murine model)
			UNC93B1 is a multitransmembrane endoplasmic
			reticulum (ER)-resident protein that controls
			homeostatic TLR7 activation by balancing TLR9 to
			TLR7 trafficking. Mice harboring a D34A mutation in
			Unc93b1 show Tlr7-dependent, systemic lethal
61027	UNC93B1	21683627	inflammation.
01027	UNC93DI	21083027	UNC93B1 participates in intracellular trafficking and
			signalling for all nucleotide-sensing Toll-like receptors
61027	UNC93B1	22164301	(TLRs).
01027	UNC33DI	22104301	Serpinb9 expression in tubular epithelial cells (TECs)
			is induced by triggering of the viral double-stranded
			RNA sensors Tlr3, Ifih1 and Ddx58. Serpinb9
			upregulation increases the threshold for granzyme B-
			mediated apoptosis in TECs and protects the kidney
			against cytotoxic insults during viral infection.
142836	Serpinb9	22167597	(Demonstrated in human)
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SERPINB9 expression in human tubular epithelial cells (TECs) is induced by triggering of the viral double-stranded RNA sensors TLR3, IFIH1 and DDX58. SERPINB9 upregulation increases the threshold for granzyme B-mediated apoptosis in TECs and protects the kidney against cytotoxic insults during viral infection.

Igf11 is a PI3K-activating ligand that increases the secretion of Il6 and Tnf in lipopolysaccharide (LPS)-stimulated mast cells, as well as attenuating the production of Il1b. 21262348

> Igf1 induces Hif1a-Tlr9 cross talk that regulates inflammatory responses in glioma cells and this regulation functions in both positive and negative feedback loops. (Demonstrated in human)

Igf1Igf1 suppresses the expression of Tlr4 to exert an anti-<br/>inflammatory effect of exercising.

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SERPINB9

Igf1

Igf1

IGF1

IGF1

IGF1

Hsf1

HSF1

HSF1

Pros1

Gas6

IGF1 is a PI3K-activating ligand that increases the secretion of IL6 and TNF in lipopolysaccharide (LPS)-stimulated mast cells, as well as attenuating the production of IL1B. (Demonstrated in murine model)

IGF1 induces HIF1A-TLR9 cross talk that regulates inflammatory responses in glioma cells and this regulation functions in both positive and negative feedback loops.

IGF1 suppresses the expression of TLR4 to exert an anti-inflammatory effect of exercising. (Demonstrated in mice)

Hsf1 is necessary to initiate host defence against Mycoplasma pneumoniae infection in the lungs through Tlr2 signalling activation.

HSF1 activated by heat shock induces the expression of ATF3, a negative regulator of IL-6, and ATF3 is necessary for heat-mediated suppression of IL-6, indicating a fever-mediated feedback loop consisting of HSF1 and ATF3.

HSF1 is necessary to initiate host defence against Mycoplasma pneumoniae infection in the lungs through TLR2 signalling activation. (Demonstrated in mice)

22043818 Pros1 works with Gas6 to synergistically suppress the basal and TLR-triggered production of inflammatory cytokines in macrophages.

PROS1 works with GAS6 to synergistically suppress<br/>the basal and TLR-triggered production of<br/>inflammatory cytokines in macrophages.PROS122043818(Demonstrated in mice)

Gas6 works with Pros1 to synergistically suppress the basal and TLR-triggered production of inflammatory 22043818 cytokines in macrophages.

53634	GAS6	11948660	GAS6 is a ligand for all members of the TAM receptor tyrosine kinase family, each of which has different affinities to GAS6.
55054	UASU	11948000	GAS6 signals through the AXL receptor and the PI3- kinase/Akt1 survival pathway, protecting
53634	GAS6	16723520	oligodendrocytes from growth factor withdrawal and TNF-alpha-mediated cell death. GAS6 works with PROS1 to synergistically suppress
53634	GAS6	22043818	the basal and TLR-triggered production of inflammatory cytokines in macrophages. (Demonstrated in mice) Cd300lf inhibits Myd88 and/or TRIF-mediated TLR
214212	Cd300lf	22043923	signalling pathway through the dual activation of Ptpn6 and Ptpn11. (Demonstrated in human) CD300LF inhibits MYD88 and/or TRIF-mediated
67452	CD300LF	22043923	TLR signalling pathway through the dual activation of PTPN6 and PTPN11. Cd300a inhibits Myd88 and/or TRIF-mediated TLR
214150	Cd300a	22043923	signalling pathway through activation of Ptpn6. (Demonstrated in human)
67271	CD300A	22043923	CD300A inhibits MYD88 and/or TRIF-mediated TLR signalling pathway through activation of PTPN6.
128168	Rictor	22045807	Rictor reduces Tlr4-mediated inflammation by regulating the cellular localization of Foxo1. RICTOR reduces TLR4-mediated inflammation by
17688	RICTOR	22045807	regulating the cellular localization of FOXO1. (Demonstrated in mice) Ccl17 is required for the induction of intestinal
			inflammation in mice. Ccl17 has an autocrine effect on dendritic cells that promotes production of
182659	Ccl17	22057112	inflammatory cytokines and activation of Th1 and Th17 cells and reduces expansion of Treg cells. IL4 (Interleukin-4) induces CCL17 expression via two
33236	CCL17	18191727	STAT6 motifs in the proximal promoter and a distal tandem STAT6 element
			CCL17 is required for the induction of intestinal inflammation in mice. CCL17 has an autocrine effect on dendritic cells that promotes production of inflammatory cytokines and activation of Th1 and Th17 cells and reduces expansion of Treg cells.
33236	CCL17	22057112	(Demonstrated in mice) Ifit2 expression imparts anti-viral immunity to restrict
159756	Ifit2	22065572	West Nile virus infection and control viral pathogenesis. IFIT2 belongs to type 1 interferon response genes and is highly induced after stimulation with LPS and
81895	IFIT2	19108715	selectively affects LPS induced protein expression by regulation at different post-transcriptional levels. IFIT2 expression imparts anti-viral immunity to restrict
81895	IFIT2	22065572	West Nile virus infection and control viral pathogenesis. (Demonstrated in mice)

			Yyl negatively regulates TLR3-induced expression of IFN-beta and acts downstream of TLR3 to limit the
170894	Yy1	22065573	level and duration of the anti-viral response. (Demonstrated in human) YY1 negatively regulates TLR3-induced expression of
20038	YY1	22065573	IFN-beta and acts downstream of TLR3 to limit the level and duration of anti-viral response. mmu-mir-10a expression is negatively regulated by intestinal microbiota, which contributes to the
223693	mmu-mir-10a	22068236	<ul><li>maintenance of intestinal homeostasis by targeting Il12b.</li><li>MIR10A expression is negatively regulated by intestinal microbiota, which contributes to the</li></ul>
127395	MIR10A	22068236	maintenance of intestinal homeostasis by targeting IL12B. (Demonstrated in mice)
35412	MRGPRX2	22069323	MRGPRX2 is a G protein-coupled receptor (GPCR) for the host defence peptide LL-37 in mast cells. IL33, a member of the IL-1-related cytokines, is the first factor shown to modulate tryptase expression in mast cells (MCs). Synovial fibroblasts promote the
48350	IL33	20427273	expression and granule accumulation of tryptase via IL33 and its receptor ST-2 (IL1RL1). IL33 attenuates sepsis by enhancing neutrophil influx to the site of infection by preventing the down- regulation of CXCR2 and reverses the TLR4-induced reduction of CXCR2 expression in neutrophils via the
48350	IL33	20473304	inhibition of expression of G protein-coupled receptor kinase-2 (GRK2). IL33 is a crucial amplifier of mucosal and systemic innate, rather than acquired, immune responses, where it is essential for manifestation of T cell-independent protease allergen-induced airway inflammation as well as OVA-induced allergic topical airway inflammation,
48350	IL33	20937871	<ul><li>without affecting acquisition of antigen-specific memory T cells.</li><li>IL33 is produced in alveolar macrophages that have been infected with Influenza A virus. The IL33-IL13</li></ul>
48350	IL33	21623379	signalling axis is required for airway hyper-reactivity in asthma. (Demonstrated in murine model) Innate lymphoid cells responding to IL33 mediate
48350	IL33	22119406	airway hyperreactivity independently of adaptive immunity. (Demonstrated in mice) Rgs2 negatively regulates Nox1 expression and
197102	Rgs2	22120521	consequently inhibits reactive oxygen species production in TLR-mediate innate immune responses. RGS2 negatively regulates NOX1 expression and consequently inhibits reactive oxygen species
105451	RGS2	22120521	production in TLR-mediated innate immune responses. (Demonstrated in mice)

			Nox1 is essential for reactive oxygen species production in the TLR-mediated innate immune response. Nox1 expression is enhanced by Tlr2
170699	Nox1	22120521	signalling through the Jak1/3-Stat3 pathway and is negatively regulated by Rgs2. NOX1 activity is regulated through MAP kinase (MAPK), protein kinase C (PKC), and protein kinase A (PKA)-dependent phosphorylation on Ser-282 and Ser-
79287	NOX1	20110267	172 of NOXA1. NOX1 is highly expressed in the colon epithelium and
79287	NOX1	18511861	can be induced by LPS or IFN-gamma. NOX1 is essential for reactive oxygen species production in the TLR-mediated innate immune response. NOX1 expression is enhanced by TLR2 signalling through the JAK1/3-STAT3 pathway and is
79287	NOX1	22120521	negatively regulated by RGS2. (Demonstrated in mice) Ppp3r1 plays an important role in activation of peritoneal macrophages. Ppp3r1 induces Tnfsf10 (TRAIL) gene expression in peritoneal macrophages in
151772	Ppp3r1	22116828	an Itgam-dependent manner. PPP3R1 plays an important role in activation of peritoneal macrophages. PPP3R1 induces TNFSF10 (TRAIL) gene expression in peritoneal macrophages in
281607	PPP3R1	22116828	an ITGAM-dependent manner. (Demonstrated in mice) PCBP2 is a negative regulator in MAVS-mediated antiviral signalling that recruits the HECT domain- containing E3 ligase ITCH (AIP4) to polyubiquitinate
36771	PCBP2	19881509	and degrade MAVS. PCBP2 synergizes with PCBP1 in MAVS inhibition but PCBP2 shows low basal expression with rapid induction after infection while PCBP1 is stably and
36771	PCBP2	22105485	abundantly expressed. Pcbp1 is critical in regulating Mavs degradation for both fine-tuning antiviral immunity and preventing
164861	Pcbp1	22105485	inflammation. (Demonstrated in human) PCBP1 is critical in regulating MAVS degradation for
55131	PCBP1	22105485	both fine-tuning antiviral immunity and preventing inflammation. Tpr53 serves as a host antiviral factor and enhances
191036	Trp53	22105999	both the innate and adaptive immune responses to influenza A virus. Fance is involved in lipopolysaccharide (LPS)-induced peritoneal macrophage inflammatory response. Fance null mice had impaired monocyte/macrophage
160833	Fance	22106009	trafficking and cytoskeletal rearrangements following LPS treatment. FANCC is involved in lipopolysaccharide (LPS)- induced peritoneal macrophage inflammatory response. Fancc null mice had impaired monocyte/macrophage trafficking and cytoskeletal rearrangements following
77041	FANCC	22106009	LPS treatment. (Demonstrated in mice)

7897	APOBEC3B	22108670	Deletion of the APOBEC3B gene strongly impacts susceptibility to Plasmodium falciparum malaria. DEFB1 is ubiquitiously expressed by all human epithelial cells, and upon reduction of the disulfide- bridges DEFB1 becomes a potent antimicrobial peptide against opportunistic pathogenic fungi and anaerobic Gram+ bacteria.
5986	DEFB1	21248850	DEFB1 is upregulated in plasmacytoid dendritic cells and monocytes during viral challenge. Defb1-deficient mice infected with mouse-adapted HK18 (influenza) lost weight earlier and died sooner than WT mice, suggesting that Defb1 plays a role in early innate immune responses against influenza in vivo. However, lung virus titers were equal between the two mouse strains, indicating that the mechanism is not related to viral replication.
5986	DEFB1	21551252	DEFB1 is a component of platelets that displays classic antimicrobial activity and signals polymorphonuclear
5986	DEFB1	22102811	leukocytes to extrude DNA lattices that capture and kill bacteria. Pglyrp3 is a member of the Peptidoglycan Recognition Proteins (PGRP) family and recognizes peptidoglycan,
168189	Pglyrp3	21439073	a structural component of bacterial cell walls, as a part of innate immune response against infections. Pglyrp3 binds to Gram-positive bacterial wall and activates a protein-sensing two-component system to induce bacterial death. PLGYRP1-mediated activation results in membrane depolarization and cessation of peptidoglycan, protein, and RNA/DNA synthesis, as well as the production of hydroxyl radicals. (Demonstrated in human)
168189	Pglyrp3	21602801	Pglyrp3, when stimulated with peptidoglycan, has an anti-inflammatory effect on intestinal epithelial cells. PGLYRP3 knocking down enhanced the expression of PGN-induced inflammatory cytokines. (Demonstrated
168189	Pglyrp3	22099350	PGLYRP3 is a secreted innate immunity protein that is expressed on body surfaces, mucous membranes, and in secretions (saliva, sweat) and is conserved from insects to mammals, it recognizes bacterial peptidoglycan, and functions in antibacterial immunity and inflammation.
102636	PGLYRP3	20418257	PGLYRP3 is a member of the Peptidoglycan Recognition Proteins (PGRP) family and recognizes peptidoglycan, a structural component of bacterial cell walls, as a part of innate immune response against
102636	PGLYRP3	21439073	infections.

PGLYRP3 binds to Gram-positive bacterial wall and activates a protein-sensing two-component system to induce bacterial death. PLGYRP3-mediated activation results in membrane depolarization and cessation of peptidoglycan, protein, and RNA/DNA synthesis, as well as the production of hydroxyl radicals.

PGLYRP3, when stimulated with peptidoglycan, has an anti-inflammatory effect on intestinal epithelial cells. PGLYRP3 knocking down enhanced the expression of PGN-induced inflammatory cytokines.

The tumour suppressor Cdkn2a plays a role in innate immunity as a regulator of inflammatory cell signalling. Cdkn2a null mice were unable to trigger a proper inflammatory response in experimental peritonitis or in induced edema and were resistant to LPS-induced endotoxic shock.

The tumour suppressor CDKN2A plays a role in innate immunity as a regulator of inflammatory cell signalling. Cdkn2a null mice were unable to trigger a proper inflammatory response in experimental peritonitis or in induced edema and were resistant to LPS-induced endotoxic shock. (Demonstrated in mice) Dcn (Decorin) is an extracellular matrix proteoglycan that stimulates proinflammatory Pdcd4 and decreases the abundance of mmu-mir-21, boosting inflammatory activity in sepsis and suppressing tumour growth.

DCN (Decorin) is an extracellular matrix proteoglycan that stimulates proinflammatory PDCD4 and decreases the abundance of MIR21, boosting inflammatory activity in sepsis and suppressing tumour growth.

MIR125B2 may enhance type I IFN expression by suppressing EIF4EBP1 protein expression in airway epithelial cells, which potentially contributes to mucosal eosinophilia in eosinophilic chronic rhinosinusitis with nasal polyps (CRSwNP).

MIR125B1 may enhance type I IFN expression by suppressing EIF4EBP1 protein expression in airway epithelial cells, which potentially contributes to mucosal eosinophilia in eosinophilic chronic rhinosinusitis with nasal polyps (CRSwNP).

Bst2 is a broad spectrum effector of the innate immune response to viral infection that is antagonized by the human immunodeficiency virus (HIV) Vpu protein to evade innate immune system detection. (Demonstrated in human)

Dlk1 is a Notch ligand that plays a critical role in the development of anti-viral immunity. Dlk1 expression in macrophages specifically regulates IFN-gamma levels from CD4(+) and CD8(+)T cells in response to influenza A H1N1 virus infection.

102636 PGLYRP3 21602801

22099350

22087031

22071331

22072710

102636 PGLYRP3

- 162905 Cdkn2a 22095712 54547 CDKN2A 22095712
- 50872 DCN 22087031

Dcn

187478

127195

171234

- 126493 MIR125B2 22071331

Dlk1

MIR125B1

- 165924 Bst2

			DLK1 is a Notch ligand that plays a critical role in the development of anti-viral immunity. DLK1 expression in macrophages specifically regulates IFN-gamma levels from CD4(+) and CD8(+)T cells in response to influenza A H1N1 virus infection. (Demonstrated in
20347	DLK1	22072963	mituenza A mitur vitus infection: (Demonstrated in mice) Tnip1 interacts with polyubiquitin to limit the activation of Tlr-Myd88 signalling pathway and prevents autoimmunity.
174207	Tnip1	21606507	Tnip1 is an essential anti-inflammatory component of
174207	Tnip1	22011580	TLR-signalling pathways that controls Cebpb activity. Tnip1 null mice exhibit progressive, lupus-like inflammatory disease. TNIP1 interacts with polyubiquitin to limit the
			activation of TLR-MYD88 signalling pathway and prevents autoimmunity. (Demonstrated in murine model)
54098	TNIP1	21606507	TNIP1 is an essential anti-inflammatory component of
54098	TNIP1	22011580	TLR-signalling pathways that controls CEBPB activity. Tnip1 null mice exhibit progressive, lupus- like inflammatory disease. (Demonstrated in mice) Rac2 is modified by Escherichia coli protein CNF1, which then interacts with the innate immune adaptors
159514	Rac2	22018470	Ripk1 and Ripk2 to trigger an anti-bacterial immune response. This response was protective and increased the ability of the host to restrict pathogen growth, thus defining a mechanism of effector-triggered immunity that contributes to how metazoans defend against microbes with pathogenic potential. (Demonstrated in human) RAC2 is modified by Escherichia coli protein CNF1, which then interacts with the innate immune adaptors RIPK1 and RIPK2 to trigger an anti-bacterial immune response. This response was protective and increased the ability of the host to restrict pathogen growth, thus
			defining a mechanism of effector-triggered immunity that contributes to how metazoans defend against
6499	RAC2	22018470	microbes with pathogenic potential. Pklr is recruited by hepatitis C virus early in infection
162642	Pklr	22022264	as a sensor to trigger the induction of Irf3-dependent genes. (Demonstrated in human) PKLR is recruited by hepatitis C virus early in
103234	PKLR	22022264	infection as a sensor to trigger the induction of IRF3- dependent genes. Senp2 acts as a negative regulator of virus-triggered
147701	Senp2	22028379	IFN-beta induction by deSUMOylating Irf3 and conditioning it for ubiquitination and degradation.

			SENP2 acts as a negative regulator of virus-triggered IFN-beta induction by deSUMOylating IRF3 and
68570	SENP2	22028379	conditioning it for ubiquitination and degradation. (Demonstrated in mice)
			Signal transducer and activator of transcription 6 (STAT6) is a transcription factor that mediates IL-13
42122	STAT6	17971840	cytokine signaling. STAT6 plays a role in the interleukin-4 (IL-4)-
42122	STAT6	16433852	mediated modulation of TLR4 expression and the responsiveness of cells to lipopolysaccharide (LPS). STAT6 is phosphorylated upon viral infection and
42122	STAT6	22000020	translocates to the nucleus to induce genes responsible for immune cell homing. (Demonstrated in mice) Fadd interacts with Trim21 to negatively regulate the late Ifna pathway in response to viral infections. Fadd enhances Trim21 ubiquitin ligase activity to repress Ifna activation in SeV infected cells. In addition, Fadd with Trim21 can ubiquitinate Irf7 and change its phosphorylation state, consequently interfering with
212910	Fadd	21183682	the activity of Traf6.
212910	Fadd	21804564	Fadd preserves epithelial barrier integrity and antibacterial defence, maintains homeostasis and prevents chronic intestinal inflammation. Upon cleavage, Fadd oligomerizes and activates the
212910	Fadd	21979465	caspase cascade to induce killing of tumour cells and intracellular pathogens by innate effector natural killer cells. (Demonstrated in human) Fadd deficiency induces severe inflammatory skin
212910	Fadd	22000287	<ul><li>lesions in mice, revealing a protective role for Fadd in epidermal keratinocytes.</li><li>FADD is an adaptor protein involved in death receptormediated apoptosis and a physiological negative</li></ul>
62144	FADD	17785432	regulator of IRAK1/MyD88-dependent responses in innate immune signalling. FADD is part of the viral product dsRNA-triggered death inducing signalling complexes (dsRNA-DISCs)
62144	FADD	15711932	containing TRADD and CASP8. FADD interacts with TRIM21 to negatively regulate the late IFNA pathway in response to viral infections. FADD enhances TRIM21 ubiquitin ligase activity to repress IFNA activation in SeV infected cells. In addition, FADD with TRIM21 can ubiquitinate IRF7 and change its phosphorylation state, consequently interfering with the activity of TRAF6.
62144	FADD	21183682	FADD preserves epithelial barrier integrity and
62144	FADD	21804564	antibacterial defence, maintains homeostasis and prevents chronic intestinal inflammation. (Demonstrated in mouse)

			Upon cleavage, FADD oligomerizes and activates the caspase cascade to induce killing of tumour cells and intracellular pathogens by innate effector natural killer
62144	FADD	21979465	cells. FADD deficiency induces severe inflammatory skin
62144	FADD	22000287	lesions in mice, revealing a protective role for FADD in epidermal keratinocytes. (Demonstrated in mice) Pla2g4a and its metabolite lipid mediators induce autophagy in macrophages and monocytes. This
197265	Pla2g4a	22003202	autophagy is Atg5 dependent and independent of changes in mTOR or autophagic flux. PLA2G4A and its metabolite lipid mediators induce autophagy in macrophages and monocytes. This autophagy is ATG5 dependent and independent of
105429	PLA2G4A	22003202	<ul> <li>changes in mTOR or autophagic flux. (Demonstrated in mice)</li> <li>P2rx7 activation in lipopolysaccharide (LPS)-primed myeloid cells results in secretion of pro-inflammatory cytokines II1b and II18. In addition, P2rx7 functions in</li> </ul>
198410	P2rx7	21988719	the recognition and phagocytosis of non-opsonized bacteria and apoptotic cells. (Demonstrated in human) Plaur is required for optimal TLR2-induced neutrophil
154827	Plaur	21998707	activation.
55442	PLAUR	21998707	PLAUR is required for optimal TLR2-induced neutrophil activation. (Demonstrated in mice) Gzmm cleaves Fadd to potentiate the killing efficacy of
170467	Gzmm	21979465	innate effector natural killer cells against tumor cells and intracellular pathogens. (Demonstrated in human) GZMM (granzyme M) cleaves BIRC5 and this triggers
12497	GZMM	20406824	degradation of the BIRC5-XIAP complex to free caspase activity, leading to cytolysis of target cells. GZMM cleaves FADD to potentiate the killing efficacy of innate effector natural killer cells against tumor cells
12497	GZMM	21979465	and intracellular pathogens. SELK is a novel target for calpain protease and its cleavage is regulated by the calpain/calpastatin system during Toll-like receptor (TLR)-induced activation of
40608	SELK	21849499	macrophages. (Demonstrated in murine model) Aire transcriptionally upregulates Tlr1/3/8 via direct interaction with the TLR promoters. In addition, Aire also mediates the induction of Il1a, Tnf, Nos2 and IFN-
168233	Aire	21628060	alpha expression upon Tlr1 and Tlr3 stimulation. Aire participates in Dectin-1 signalling, an anticandidal innate immune signalling pathway that specifically recognizes fungal beta-glucan. Aire formes a transient complex with the known Dectin-1 pathway components phosphorylated Sykb and Card9 after receptor ligation and localizes with Clec7a (Dectin-1)
168233	Aire	21962774	at the cell membrane (Demonstrated in human)

5152	AIRE	21628060	AIRE transcriptionally upregulates TLR1/3/8 via direct interaction with the TLR promoters. In addition, AIRE also mediates the induction of IL1A, TNF, NOS2 and IFN-alpha expression upon TLR1 and TLR3 stimulation. (Demonstrated in murine model) AIRE participates in Dectin-1 signalling, an anticandidal innate immune signalling pathway that specifically recognizes fungal beta-glucan. AIRE formes a transient complex with the known Dectin-1 pathway components phosphorylated SYK and
5152	AIRE	21962774	CARD9 after receptor ligation and localizes with CLEC7A (Dectin-1) at the cell membrane. Apoh is a plasma protein that specifically interacts with
213761	Apoh	21965665	lipopolysaccharide (LPS) and Tlr4 to induce Tnf-alpha production in macrophages. (Demonstrated in human) APOH is a plasma protein that specifically interacts
65172	АРОН	21965665	with lipopolysaccharide (LPS) and TLR4 to induce TNF-alpha production in macrophages.
151805	Ppargc1b	21966468	Ppargc1b is activated in Staphylococcus aureus- mediated sepsis via the Tlr2-signalling pathway. PPARGC1B is activated in Staphylococcus aureus-
53101	PPARGC1B	21966468	mediated sepsis via the TLR2-signalling pathway. (Demonstrated in mice)
163551	Ppargc1a	21966468	Ppargc1a is activated in Staphylococcus aureus- mediated sepsis via the Tlr2-signalling pathway. Mapkapk2 phosphorylation is suppressed by lipomannan derived from virulent Mycobacterium tuberculosis (M.tb), resulting in the potent suppression of Tnf biosynthesis and allowing M.tb to subvert host
190728	Mapkapk2	21969554	<ul> <li>immunity and potentially to increase its virulence.</li> <li>(Demonstrated in human)</li> <li>MAPKAPK2 phosphorylation is suppressed by lipomannan derived from virulent Mycobacterium tuberculosis (M.tb), resulting in the potent suppression</li> </ul>
106265	МАРКАРК2	21969554	of TNF biosynthesis and allowing M.tb to subvert host immunity and potentially to increase its virulence. Zfpm2 inhibits the transcription of hepcidin
137408	Zfpm2	21971825	antimicrobial peptide by suppressing the GATA4 and GATA6 transcription factors. (Demonstrated in human) ZFPM2 inhibits the transcription of hepcidin
32130	ZFPM2	21971825	antimicrobial peptide by suppressing the GATA4 and GATA6 transcription factors. Zfpm1 inhibits the transcription of hepcidin
197075	Zfpm1	21971825	antimicrobial peptide by suppressing the GATA4 and GATA6 transcription factors. (Demonstrated in human) ZFPM1 inhibits the transcription of hepcidin
46137	ZFPM1	21971825	antimicrobial peptide by suppressing the GATA4 and GATA6 transcription factors.

			Gata6 activates the transcription of hepcidin antimicrobial peptide in hepatocytes. Friend of GATA (FOG)-proteins ZFPM1 and ZFPM2 moderate this transcription by suppressing the GATA transactivation
128584	Gata6	21971825	of hepcidin promoter. (Demonstrated in human) GATA6 activates the transcription of hepcidin antimicrobial peptide in hepatocytes Friend of GATA (FOG)-proteins ZFPM1 and ZFPM2 moderate this transcription by suppressing the GATA transactivation
1469	GATA6	21971825	of hepcidin promoter. Gata4 activates the transcription of hepcidin antimicrobial peptide in hepatocytes. Friend of GATA (FOG)-proteins ZFPM1 and ZFPM2 moderate this transcription by suppressing the GATA transactivation
172452	Gata4	21971825	of hepcidin promoter. (Demonstrated in human) GATA4 activates the transcription of hepcidin antimicrobial peptide in hepatocytes. Friend of GATA (FOG) proteins, ZFPM1 and ZFPM2, moderate this transcription by suppressing the GATA transactivation
7434	GATA4	21971825	of hepcidin promoter. Tyk2 functions at the molecular interface between innate immunity and cellular metabolism and is involved in the regulation of lipid and carbohydrate
139044	Tyk2	21787891	metabolism in macrophages stimulated with poly(I:C). Tyk2 is necessary for Socs1-mediated suppression of
139044	Tyk2	21757742	Type I IFN signalling. TYK2 is directly involved in IFN-alpha signalling for the induction and translocation of Daxx nuclear protein, which may result in B lymphocyte growth
27182	TYK2	12391177	arrest and/or apoptosis. TYK2 modulates the relationship between immunity and metabolism where it is essential for the full LPS response, its function mainly being required for baseline expression and not LPS-induced upregulation of IFN-inducible genes, as well as its critical role in the downregulation of metabolic genes upon immune challenge, in particular genes involved in lipid metabolism.
27182	TYK2	20338026	TYK2 functions at the molecular interface between innate immunity and cellular metabolism and is involved in the regulation of lipid and carbohydrate metabolism in macrophages stimulated with poly(I:C).
27182	TYK2	21787891	(Demonstrated in mouse)
27182	TYK2	21757742	TYK2 is necessary for SOCS1-mediated suppression of Type I IFN signalling. (Demonstrated in mice) The Glucocorticoid receptor, Nr3c1 exerts anti- inflammatory action in part by antagonizing pro- inflammatory transcription factors such as Rela of the
142259	Nr3c1	21750107	NFkB complex. (Demonstrated in human)

			Nr3c1, the glucocorticoid receptor, is necessary for glucocorticoid-mediated induction of Nlrp3 and the subsequent formation of inflammasomes, demonstrating a novel role for glucocorticoids in
142259	Nr3c1	21940629	sensitizing the initial inflammatory response by the innate immune system. The Glucocorticoid receptor, NR3C1, exerts anti- inflammatory action in part by antagonizing pro- inflammatory transcription factors such as RELA of the
51637	NR3C1	21750107	NFkB complex. NR3C1, the glucocorticoid receptor, is necessary for glucocorticoid-mediated induction of NLRP3 and the subsequent formation of inflammasomes, demonstrating a novel role for glucocorticoids in sensitizing the initial inflammatory response by the
51637	NR3C1	21940629	innate immune system. (Demonstrated in mice) Stub1 facilitates the formation of a TLR signalling complex by recruiting, ubiquitinating, and activating Src and Prkcz. Knockdown of Stub1 inhibits Tlr4- and
153401	Stub1	21911421	Ste and Prkez. Knockdown of Stubi inhibits 114- and Tlr9-signalling pathways. STUB1 facilitates the formation of a TLR signalling complex by recruiting, ubiquitinating, and activating Src and PRKCZ. Knockdown of Stub1 in mice inhibits Tlr4- and Tlr9-signalling pathways. (Demonstrated in
8030	STUB1	21911421	mice) Serpinel is a critical mediator that controls the development of the early lung inflammation induced by
204786	Serpine1	21768189	Pseudomonas aeruginosa infections. Serpinel plays a critical role in early host defense response against Haemophilus influenzae infection. Serpinel knockout mice show reduced bacterial
204786	Serpine1	21945446	clearance and prolonged pneumonia. SERPINE1 is a critical mediator of the pulmonary host response that controls the development of the early lung inflammation induced by Pseudomonas
32959	SERPINE1	21768189	aeruginosa infections. (Demonstrated in mouse) SERPINE1 plays a critical role in early host defense response against Haemophilus influenzae infection. Serpine1 knockout mice show reduced bacterial
32959	SERPINE1	21945446	clearance and prolonged pneumonia. (Demonstrated in mice) Tank plays a key role in the cross-talk between the IKK-related and the canonical IKK kinases, a
173742	Tank	21949249	mechanism required to limit the strength of TLR- signalling and prevent autoimmunity. TANK is involved in interferon responses and is a negative regulator of pro-inflammatory cytokine
73552	TANK	19668221	production induced by TLR signalling.

TANK plays a key role in the cross-talk between the IKK-related and the canonical IKK kinases, a mechanism required to limit the strength of TLR-signalling and prevent autoimmunity. (Demonstrated in mice)

72550		01040040	signalling and prevent autoimmunity. (Demonstrated in
73552	TANK	21949249	mice)
			NAIP is important for CASP1 activation and IL-1
26953	NAIP	18280719	processing.
			NAIP specifically recognizes the Chromobacterium
			violaceum type III secretion system protein, Cprl, and
26953	NAIP	21918512	activates the NLRC4 inflammasome in macrophages.
			Naip2 functions as a specific inflammasome receptor
			for the type III secretion system rod component of
262522	Naip2	21918512	Salmonella and Burkholderia bacterial species.
	1		Trem2 inhibits the induction of inflammatory
			cytokines and type I IFN production in TLR-stimulated
189871	Trem2	21956652	dendritic cells.
10/0/1	1101112	21)50052	TREM2 inhibits the induction of inflammatory
			•
0(5((		2105((52	cytokines and type I IFN production in TLR-stimulated
86566	TREM2	21956652	dendritic cells. (Demonstrated in mice)
			APOBEC3G is an innate intracellular HIV restriction
402349	APOBEC3G	21734563	factor that is upregulated by type I interferons.
			APOBEC3G, an intrinsic antiviral factor, promotes the
			recognition of human immunodeficiency virus (HIV)
			through the activation of the DNA-damage response
			pathway and the expression of natural killer cell-
402349	APOBEC3G	21874023	activating ligands on HIV-infected cells.
			Ddx41 is a helicase responsible for sensing intracellular
			DNA in myeloid dendritic cells. Ddx41 expression
			knockdown blocked the induction of type I interferon
156279	Ddx41	21892174	and cytokine responses to DNA and DNA viruses.
			DDX41 is a helicase responsible for sensing
			intracellular DNA in myeloid dendritic cells. DDX41
			expression knockdown blocked the induction of type I
			interferon and cytokine responses to DNA and DNA
60413	DDX41	21892174	viruses. (Demonstrated in mice)
			Tax1bp1 phosphorylation by Chuk is pivotal for the
			proinflammatory cytokine-dependent assembly of the
			A20 ubiquitin-editing complex to limit inflammatory
148305	Tax1bp1	21765415	gene activation.
140505	Taxiopi	21703413	TAX1BP1 is a nuclear receptor co-activator that forms
			a complex with the glucocorticoid receptor (NR3C1).
			TAX1BP1 can bind Tax, a human T-cell leukemia
			virus type 1 oncoprotein, directly and this induces the
			dissociation of TAX1BP1 from the glucocorticoid
10(00		17000140	receptor-containing protein complex, and represses the
10692	TAX1BP1	17283140	co-activator function of TAX1BP1.
			TAX1BP1 and TNFAIP3 (A20) inhibit antiviral
10.00	m	00001010	signalling by targeting TBK1/IKKi kinases and
10692	TAX1BP1	20304918	disrupting the TRAF3-TBK1-IKKi signalling complex.

TAX1BP1 phosphorylation by CHUK is pivotal for the proinflammatory cytokine-dependent assembly of the A20 ubiquitin-editing complex to limit inflammatory gene activation. (Demonstrated in mouse)

Neu1 and Mmp9 cross-talk in alliance with Tlr4 on the cell surface is a novel membrane sialidase-controlling mechanism that depends on ligand binding to its Tolllike receptor (TLR) to induce Neu1 activity, to influence receptor desialylation and subsequently to induce TLR receptor activation and the production of nitric oxide and pro-inflammatory cytokines in dendritic and macrophage cells.

NEU1 and MMP9 cross-talk in alliance with TLR4 on the cell surface is a novel membrane sialidasecontrolling mechanism that depends on ligand binding to its Toll-like receptor (TLR) to induce NEU1 activity, to influence receptor desialylation and subsequently to induce TLR receptor activation and the production of nitric oxide and pro-inflammatory cytokines in dendritic and macrophage cells. (Demonstrated in murine model)

Aqp3 is an aquaporin that functions in macrophage immunity by a cellular mechanism involving facilitated water and glycerol transport, and consequent phagocytic and migration activity. Aqp3, therefore, is a novel therapeutic target in modulating the innate immune response in various infectious and inflammatory conditions.

AQP3 is an aquaporin that functions in macrophage immunity by a cellular mechanism involving facilitated water and glycerol transport, and consequent phagocytic and migration activity. AQP3, therefore, is a novel therapeutic target in modulating the innate immune response in various infectious and inflammatory conditions.

Elf1 transcription factor negatively regulates Tollip, a negative regulator of Toll-like receptor (TLR) signalling, in response to O-linked Nacetylglucosamine (O-GlcNAc) modification. In intestinal epithelial cells insufficient O-GlcNAc modification prevents Elf1-mediated transcriptional repression and thereby suppresses TLR signalling via upregulated Tollip gene expression. (Demonstrated in human)

ELF1 transcription factor negatively regulates TOLLIP, a negative regulator of Toll-like receptor (TLR) signalling, in response to O-linked Nacetylglucosamine (O-GlcNAc) modification. In intestinal epithelial cells, insufficient O-GlcNAc modification prevents ELF1-mediated transcriptional repression and thereby suppresses TLR signalling via upregulated TOLLIP gene expression.

10692TAX1BP121765415gene a<br/>Neu1Neu1cell su<br/>mecha<br/>like r<br/>influer<br/>induce<br/>nitric175208Neu121873432dendri<br/>NEU1<br/>the c<br/>contro<br/>to its T<br/>to infl<br/>induce<br/>nitric175398NEU121873432murine

137002 Aqp3 21865318

- 57960 AQP3 21865318

21867680

21867680

Elf1

ELF1

182872

26868

155582	Cd8a	21867928	Cd8a-positive dendritic cells play a critical role in cross-presentation of antigens during intracellular pathogen infections, specifically by activating an innate immune response through Il12 production. CD8A-positive dendritic cells play a critical role in cross-presentation of antigens during intracellular pathogen infections, specifically by activating an innate
59987	CD8A	21867928	<ul><li>immune response through IL12 production.</li><li>(Demonstrated in murine model)</li><li>Muc1 controls the inflammatory response in airway</li><li>epithelial cells during nontypeable Haemophilus</li><li>influenzae infection, mainly through suppression of</li></ul>
163463	Muc1	21868711	Tlr2 signalling and decreased Il8 production. MUC1 regulates innate immune responses of dendritic cells (DC) where deletion of MUC1 promotes a heightened functional response of DC in response to
103122	MUC1	20375631	TLR4 and TLR5 signalling pathways. MUC1 controls the inflammatory response in airway epithelial cells during nontypeable Haemophilus influenzae infection, mainly through suppression of
103122	MUC1	21868711	<ul><li>TLR2 signalling and decreased IL8 production.</li><li>(Demonstrated in murine model)</li><li>Il1rl2 binds members of the interleukin (IL)-1 family of cytokines that includes Il1f6, Il1f8 and Il1f9. Binding of these ligands to Il1rl2 plays a critical role in the interface between innate and adaptive immunity,</li></ul>
148344	Il1rl2	21860022	<ul><li>leading to the stimulation of dendritic and T helper cell responses.</li><li>IL1RL2 binds members of the interleukin (IL)-1 family of cytokines that includes IL1F6, IL1F8 and IL1F9.</li><li>Binding of these ligands to IL1RL2 plays a critical role in the interface between innate and adaptive immunity,</li></ul>
64113	IL1RL2	21860022	leading to the stimulation of dendritic and T helper cell responses. (Demonstrated in murine model) Rpl19 inhibits Irf3 activation and Cxcl10 production to facilitate viral multiplication in cells that express Tlr3 in endosomes, and Rpl19 inhibits viral multiplication
210447	Rpl19	21860608	in cells bearing Tlr3 on their cell membrane. (Demonstrated in human) RPL19 inhibits IRF3 activation and CXCL10 production to facilitate viral multiplication in cells that express TLR3 in endosomes, and RPL19 inhibits viral
45490	RPL19	21860608	<ul><li>multiplication in cells bearing TLR3 on their cell membrane.</li><li>Selk is a novel target for calpain protease and its cleavage is regulated by the calpain/calpastatin system during Toll-like receptor (TLR)-induced activation of</li></ul>
142010	Selk	21849499	macrophages.

BCL2 is a multifunctional regulator of cell survival that inhibits the innate immune response during early stages of pathogenesis. Muscle-specific expression of BCL2 in Lama2-deficient mice resulted in the inhibition of TLR4, TLR6, TLR7, TLR8 and TLR9 induction, leading to reduced infiltration of the principal death eosinophils. effector cells. (Demonstrated in murine model)

4354 BCL2 21850221 The Il4ra signalling pathway contributes to the heightened susceptibility of mice co-infected with Mycobacterium tuberculosis and the intestinal 209275 21825018 Il4ra helminth, Nippostrongylus brasiliensis IL4R is utilized by IL13 to induce phosphorylation and activation of JAK2 in human colon carcinoma cell 21808 8609418 IL4R lines. The IL4R signalling pathway contributes to the heightened susceptibility of mice co-infected with Mycobacterium tuberculosis and the intestinal helminth, Nippostrongylus brasiliensis. (Demonstrated 21808 IL4R 21825018 in mice)

Sftpd binds Mycobacterium to avium lipoarabinomannan and resulting in the agglutination of the pathogen. The presence of Sftpd promotes the phagocytosis of M. avium by macrophages. (Demonstrated in human)

SFTPD inhibits lipopolysaccharide (LPS)-induced inflammatory cell responses by altering LPS binding to its receptors where it binds to a complex of TLR4/LY96 high (MD-2) with affinity and significantly reduces LY96 binding to both serotypes of LPS.

SFTPD, upon S-nitrosylation, controls inflammatory function by acting as a chemoattractant for macrophages and inducing p38 MAPK phosphorylation.

SFTPD is a carbohydrate pattern recognition molecule of innate immunity that significantly enhances phagocytosis and killing of Aspergillus fumigatus, a pathogenic fungus, by neutrophils and macrophages.

SFTPD can attenuate bacterial and viral infection and inflammation by acting as an opsonin and by regulating innate immune cell functions.

SFTPD binds to Mycobacterium avium lipoarabinomannan and resulting in the agglutination of the pathogen. The presence of SFTPD promotes the 21821801 phagocytosis of M. avium by macrophages.

Ifit3 triggers host antiviral responses by bridging Tbk1 to Mavs, and Ifit3 plays an important role in the 21813773 activation of Irf3. (Demonstrated in human)

- 151871 Sftpd 21821801
- 18990700 80187 SFTPD
- 80187 SFTPD 19007302
- 80187 SFTPD 20413160

20418258

80187 SFTPD

SFTPD

Ifit3

80187

159817

			mmu-mir-29b-1 suppresses immune responses to intracellular pathogens by targeting IFN-gamma mRNA. Mice infected with Listeria monocytogenes or
224213	mmu-mir-29b-1	21785411	Mycobacterium bovis bacillus Calmette-Gu à © rin (BCG) downregulated miR-29 expression. Pltp may play a pivotal role in inflammation and innate
212419	Pltp	21787334	<ul><li>immunity through its ability to accelerate the 'reverse LPS transport' pathway.</li><li>PLTP may play a pivotal role in inflammation and innate immunity through its ability to accelerate the</li></ul>
78556	PLTP	21787334	<ul><li>'reverse LPS transport' pathway. (Demonstrated in mouse)</li><li>PLUNC is a protein able to inhibit Mycoplasma pneumoniae (Mp) growth and its production following</li></ul>
66205	PLUNC	21054862	Mp infection is regulated through Toll-like receptor 2 (TLR2) signalling, as determined in mice. PLUNC is a secretory protein that exhibits antimicrobial activity against Gram-negative bacteria
66205	PLUNC	21787346	and anti-inflammatory functions in respiratory infections. Ltbr signalling in the gut lymphoid follicles regulates
189788	Ltbr	21767811	<ul><li>II22 production by innate lymphoid cells in response to mucosal pathogen challenge.</li><li>LTBR signalling in intestinal epithelial cells are bestated innate immune responses against mucosal</li></ul>
13973	LTBR	20226692	orchestrates innate immune responses against mucosal bacterial infection. LTBR signalling in the gut lymphoid follicles regulates IL22 production by innate lymphoid cells in response
13973	LTBR	21767811	to mucosal pathogen challenge. (Demonstrated in mouse) Rag1 knockout mice display deficient early inflammatory responses and reduced survival during
193190	Rag1	21746813	sepsis, which demonstrates a novel role of B cells in the early innate immune response. Rag1 knockout mice display deficient early
40149	RAG1	21746813	inflammatory responses and reduced survival during sepsis, which demonstrates a novel role of B cells in the early innate immune response. (Demonstrated in mice)
			The expression of ligand-receptor pair Tnfsf9-Tnfrsf9 on monocytes and NK cells is induced by Mycobacterium tuberculosis. Blockage of the Tnfrsf9 pathway enhanced the level of Ifng and Tnfa producing
193693	Tnfsf9	21747409	lymphocytes against the pathogen. (Demonstrated in human) The expression of ligand-receptor pair TNFSF9- TNFRSF9 on monocytes and NK cells is induced by Mycobacterium tuberculosis. Blockage of the
21688	TNFSF9	21747409	TNFRSF9 pathway enhanced the level of IFNG and TNFA producing lymphocytes against the pathogen.

			The expression of ligand-receptor pair Tnfsf9-Tnfrsf9 on monocytes and NK cells is induced by Mycobacterium tuberculosis. Blockage of the Tnfrsf9 pathway enhanced the level of Ifng and Tnfa producing
205902	Tnfrsf9	21747409	lymphocytes against the pathogen. (Demonstrated in human) The expression of the ligand-receptor pair TNFSF9- TNFRSF9 on monocytes and natural killer cells is
88268	TNFRSF9	21747409	induced by Mycobacterium tuberculosis. Blockage of the TNFRSF9 pathway enhanced the level of IFNG and TNFA producing lymphocytes against the pathogen. Snca acts as a danger-associated molecular pattern that activates the expression of TLRs to initiate the
152558	Snca	21747756	proinflammatory pathway and microglial activation. (Demonstrated in human) SNCA acts as a danger-associated molecular pattern
30077	SNCA	21747756	that activates the expression of TLRs to initiate the proinflammatory pathway and microglial activation. F11 modulates the inflammatory response of PMN leukocytes by reducing chemotaxis triggered by II8 or
152055	F11	21807745	<ul><li>fMLP, highlighting the interplay between inflammation and coagulation.</li><li>F11 modulates the inflammatory response of PMN leukocytes by reducing chemotaxis triggered by IL8 or</li></ul>
47627	F11	21807745	fMLP, highlighting the interplay between inflammation and coagulation (Demonstrated in mouse) ADAR destabalizes RNA structure by the deamination
102968	ADAR	21809195	of adenosine to inosine, and therefore is able to disrupt replication of dsRNA viruses in the host. Tnfrsf1a modulates nitric oxide production in peritoneal macrophages in response to Yersinia LPS
189844	Tnfrsfla	21802165	stimulation through the II6 and NFkB signalling pathway.
13807	TNFRSF1A	12753742	Recruitment of TNF receptor 1 to lipid rafts is essential for TNFalpha-mediated NF-kappaB activation TNFR1 mediates TRAF2 phosphorylation and
13807	TNFRSF1A	19150425	TNFR1 mediates TRAF2 phosphorylation and recruitment of the IKK complex TNFRSF1A modulates nitric oxide production in peritoneal macrophages in response to Yersinia LPS
13807	TNFRSF1A	21802165	stimulation through the IL6 and NFkB signalling pathway. (Demonstrated in mouse) Tlt2 enhances neutrophil functions such as antibacterial
189835	Treml2	21804015	activity and chemotaxis by potentiating the response to the G protein-coupled receptor-signalling pathway. TLT2 enhances neutrophil functions such as antibactorial activity and abametavia by notantiating
86610	TREML2	21804015	antibacterial activity and chemotaxis by potentiating the response to the G protein-coupled receptor- signalling pathway. Cbl, an E3 ligase, has a crucial role in regulating
155970	Cbl	21799517	dendritic cell maturation by facilitating the regulatory functions of Nfkb1.

74417	CBL	21799517	CBL, an E3 ligase, has a crucial role in regulating dendritic cell maturation by facilitating the regulatory functions of NFKB1. (Demonstrated in mouse) Hspd1 plays a dual role as an immune modulator and a biomarker, and is a target to modulate immunity for therapeutic purposes, and to monitor the immune
155162	Hspd1	21145789	response in health and disease. Hspd1 and Tlr4 mediate myocardial ischemia-activated innate immune signalling, which plays an important
155162	Hspd1	21775438	role in mediating apoptosis and inflammation during ischemia/reperfusion (I/R). HSPD1 is part of the heat shock family of proteins that
77925	HSPD1	18575269	<ul><li>have many roles in inflammation and regulation of the immune system.</li><li>HSPD1 plays a dual role as an immune modulator and a biomarker, and is a target to modulate immunity for therapeutic purposes, and to monitor the immune response in health and disease.</li></ul>
77925	HSPD1	21145789	HSPD1 and TLR4 mediate myocardial ischemia- activated innate immune signalling, which plays an important role in mediating apoptosis and inflammation during ischemia/reperfusion (I/R).
77925	HSPD1	21775438	(Demonstrated in murine model) Pml, a member of the Trim protein family, is upregulated by Type I and Type II interferons and have been found to restrict viral replication by modulating the RIG-I pathway.
171712	Pml	21131187	Pml(-/-) mice are resistant to LPS-induced septic shock as a result of an ineffective production of cytokines and chemokines, suggesting a role for PML in the innate immune Toll-like receptor (TLR)/NF-kB pathway. Pml(-/-) mice also exhibit impaired function of macrophages and are thus unable to clear pathogenic
171712	Pml	21779477	microorganisms. PML, a member of the TRIM protein family, is upregulated by Type I and Type II interferons and have been found to restrict viral replication by modulating
21462	PML	21131187	the RIG-I pathway. Pml(-/-) mice are resistant to LPS-induced septic shock as a result of an ineffective production of cytokines and chemokines, suggesting a role for PML in the innate immune Toll-like receptor (TLR)/NF-kB pathway. Pml(-/-) mice also exhibit impaired function of
21462	PML	21779477	macrophages and are thus unable to clear pathogenic microorganisms. (Demonstrated in murine model) Pin1 is necessary to mount TLR-mediated, interferon- dependent innate and adaptive immune response. Pin1 is activated by Tlr7 and Tlr9, which binds to Irak1 to
137359	Pin1	21743479	activate Irf7 which then induce type I interferons.

26087	PIN1	16699525	PIN1 negatively regulates IRF3-dependent innate antiviral response by binding to phosphorylated IRF3 for proteasome-dependent degradation. PIN1 is necessary to mount TLR-mediated, interferon- dependent innate and adaptive immune response. PIN1
26087	PIN1	21743479	is activated by TLR7 and TLR9, which binds to IRAK1 to activate IRF7 which then induce type I interferons. (Demonstrated in mouse) Hrg binds fibrinogen with high affinity and competes with thrombin for binding. This interaction may
148872	Hrg	21757718	provide a novel link between coagulation, innate immunity and inflammation. (Demonstrated in human) Notch1 signalling modulates the microglia innate response to post-ischemic brain damage inflammation. Notch1 deficient mice exhibit significantly lower levels
149571	Notch1	21737799	of activated microglia and reduced proinflammatory cytokine expression. HRG binds fibrinogen with high affinity and competes with thrombin for binding. This interaction may
69039	HRG	21757718	provide a novel link between coagulation, innate immunity and inflammation. NOTCH1 signalling modulates the microglia innate response to post-ischemic brain damage inflammation. Notch1 deficient mice exhibit significantly lower levels
92445	NOTCH1	21737799	of activated microglia and reduced proinflammatory cytokine expression. (Demonstrated in mouse) Gbp2 localizes to intracellular compartments positive
197330	Gbp2	21757726	for macroautophagy markers and Immunity-Related GTPases (IRGs) indirectly modulate Gbp2 localization. GBP2 localizes to intracellular compartments positive for macroautophagy markers and Immunity-Related GTPases (IRGs) indirectly modulate GBP2
100168	GBP2	21757726	localization. (Demonstrated in murine model) Hspa14 binds directly to Tlr4 on dendritic cell surfaces
133420	Hspa14	21730052	and induces a robust Th1 response via the MAPK and NFkB signalling pathways. HSPA14 binds directly to TLR4 on dendritic cell
55933	HSPA14	21730052	surfaces and induces a robust Th1 response via the MAPK and NFkB signalling pathways. Tgtp1 is part of the Immunity-Related GTPases (IRGs) family of proteins that are induced by interferon- gamma (Ifng) and play a crucial role in innate resistance to intracellular pathogens. Tgtp1 also
262889	Tgtp1	21757726	influences the localization of Gbp2 by modulating macroautophagy. Igtp is responsible for the IFN-gamma mediated induction of the antimicrobial defence system against Chlamydia trachomatis in mouse. Irgm1/Igtp knockout mice develop high bacterial burden post intrauterine infection, but subsequently clear the infection more
263693	Igtp	21731484	efficiently than wt mice due to a compensatory T cell response.

			Igtp is part of the Immunity-Related GTPases (IRGs) family of proteins that are induced by interferon- gamma (Ifng) and play a crucial role in innate resistance to intracellular pathogens. Igtp also
263693	Igtp	21757726	influences the localization of Gbp2 by modulating macroautophagy. Edn1 links Tlr7 inflammatory signalling to cardiac
147056	Edn1	21730058	fibrosis in autoimmune associated congenital heart block. (Demonstrated in human) Socs1 is a negative immunomodulator that is upregulated by Hepatitis C virus to deliver negative signaling to Tlr-mediated pathways controlling expression of Il12, a key cytokine linking innate and adaptive immunity.
132667	Socs1	21263070	Socs1 is targeted by endogenous and pharmacologic glucocorticoids to limit Tlr3/Tlr4-mediated Stat1 activation which results in the suppression of inflammation.
132667	Socs1	21606371	Socs1 inhibits type I interferon (IFN) signaling through an interaction with the Ifnar1 associated kinase, Tyk2, resulting in a reduced IFN response and reduced Ifnar1
132667	Socs1	21757742	surface expression. (Demonstrated in human) EDN1 links TLR7 inflammatory signalling to cardiac
61847	EDN1	21730058	fibrosis in autoimmune associated congenital heart block. CD5L (AIM) is required for obesity-associated
103716	CD5L	21730133	recruitment of inflammatory macrophages into adipose tissue. (Demonstrated in mouse)
14621	SOCS1	16451196	SOCS1 is a potent and multifaceted regulator of cytokines and cell-mediated inflammation. SOCS1 regulates the IFN but Not NF{kappa}B Pathway in TLR-Stimulated Human monocytes and
14621	SOCS1	19017994	macrophages. SOCS1 is a negative immunomodulator that is upregulated by Hepatitis C virus to deliver negative signalling to TLR-mediated pathways controlling expression of IL12, a key cytokine linking innate and adaptive immunity.
14621	SOCS1	21263070	SOCS1 is targeted by endogenous and pharmacologic glucocorticoids to limit TLR3/TLR4-mediated STAT1 activation which results in the suppression of
14621	SOCS1	21606371	inflammation. (Demonstrated in murine model) SOCS1 inhibits type I interferon (IFN) signaling through an interaction with the IFNAR1 associated
14621	SOCS1	21757742	kinase, TYK2, resulting in a reduced IFN response and reduced IFNAR1 surface expression.

180975	Nras	21757746	Nras is a small GTPase that, in response to microbial activation, mediates cholangiocyte pro-inflammatory cytokine production and induction of cholangiocyte proliferation. (Demonstrated in human) Irgm1 is responsible for the IFN-gamma mediated induction of the antimicrobial defence system against Chlamydia trachomatis in mouse. Irgm1/Igtp knockout mice develop high bacterial burden post intrauterine infection, but subsequently clear the infection more
166425	Irgm 1	21731484	efficiently than wt mice due to a compensatory T cell response. NRAS is a small GTPase that, in response to microbial activation, mediates cholangiocyte pro-inflammatory cytokine production and induction of cholangiocyte
243969	NRAS	21757746	proliferation. Aimp1 is a pleiotropic cytokine expressed in the salivary glands, small intestine and large intestine. Aimp1 not only induces the maturation and activation of bone marrow-derived dendritic cells, but also
194580	Aimpl	21711348	induces the expression of Tlr1, 2, 3 and 7. MIF is a chemokine-like inflammatory mediator that triggers leukocyte recruitment by binding to CXCR2
401910	MIF	17435771	and CXCR4. (Demonstrated in murine model) Gm16379 (Mif) is a chemokine-like inflammatory
163956	Gm16379	17435771	<ul> <li>mediator that triggers leukocyte recruitment by binding to Cxcr2 and Cxcr4.</li> <li>Duox2 plays pivotal roles in the Tlr5-dependent inflammatory response of nasal airway epithelium.</li> <li>Duox2 activation is required for flagellin-induced reactive oxygen species production, as well as the induction of mucin and MIP-2alpha in nasal epithelial</li> </ul>
202825	Duox2	21714724	cells. DUOX2 is a member of the NAD(P)H oxidase family
10041	DUOX2	19759286	and is involved in NOD2-dependent reactive oxygen species (ROS) production. DUOX2 and DUOX1 localize to the apical plasma membrane of epithelial cells in major airways, salivary glands, and the gastrointestinal tract, and provide extracellular hydrogen peroxide to lactoperoxidase to produce antimicrobial hypothiocyanite ions. Expression of dual oxidases DUOX2 and DUOX1 is
10041	DUOX2	18511861	regulated by Th1 and Th2 cytokines in human airways. DUOX2 expression is mediated by IFN-gamma via a STAT-independent signalling pathway, providing insights into a novel IFN-gamma signalling pathway with potential importance for regulation of host defence
10041	DUOX2	20381453	responses.

			DUOX2 plays pivotal roles in the TLR5-dependent inflammatory response of nasal airway epithelium. DUOX2 activation is required for flagellin-induced reactive oxygen species production, as well as the induction of mucin and MIP-2alpha in nasal epithelial
10041	DUOX2	21714724	cells. Rarres2 (Chemerin) requires C-terminal proteolytic processing by cysteine cathepsins to selectively attract a specific subset of immunoregulatory APCs, such as
145192	Rarres2	21715684	immature plasmacytoid dendritic cells. In addition, truncated Rarres2 also displays antibacterial activity against Enterobacteriaceae. (Demonstrated in human) RARRES2 (Chemerin) requires C-terminal proteolytic processing by cysteine cathepsins to selectively attract a specific subset of immunoregulatory APCs, such as immature plasmacytoid dendritic cells. In addition, truncated RARRES2 also displays antibacterial activity
47881	RARRES2	21715684	against Enterobacteriaceae.
196988	Kcnj8	21719711	Kcnj8 is an ATP-sensitive potassium channel that restricts cardiotropic RNA virus replication. Nfatc4 is a transcription factor required for the
166063	Nfatc4	21726630	regulation of the Toll-like receptor-activated innate inflammatory response in monocytes/macrophages. KCNJ8 is an ATP-sensitive potassium channel that
23052	KCNJ8	21719711	restricts cardiotropic RNA virus replication. (Demonstrated in murine model) NFATC4 is a transcription factor required for the regulation of the Toll-like receptor-activated innate
3927	NFATC4	21726630	<ul><li>inflammatory response in monocytes/macrophages.</li><li>(Demonstrated in murine model)</li><li>Nfatc3 is a transcription factor required for the</li></ul>
189544	Nfatc3	21726630	regulation of the Toll-like receptor-activated innate inflammatory response in monocytes/macrophages. NFATC3 is a transcription factor required for the regulation of the Toll-like receptor-activated innate
38028	NFATC3	21726630	inflammatory response in monocytes/macrophages. (Demonstrated in murine model) Clec1b is expressed in myeloid cells and acts as a Syk- coupled C-type lectin receptor (CLR) able to modulate
191926	Clec1b	21728173	Toll-like receptor (TLR) signaling and inflammatory responses. CLEC1B is expressed in myeloid cells and acts as a SYK-coupled C-type lectin receptor (CLR) able to
18309	CLEC1B	21728173	modulate Toll-like receptor (TLR) signaling and inflammatory responses. (Demonstrated in murine model) Dhx36 is a component of a cytosolic viral sensor and is recruited to a complex consisting of Ddx1, Ddx21 and Ticam1, which triggers type I interferon and cytokine
147779	Dhx36	21703541	response to dsRNA.

			DHX36 interacts with CpG-A and is associated with IFN-alpha production and IRF7 nuclear translocation upon CpG-A stimulation. DHX36 localizes within the
62011	DHX36	20696886	cytosol and directly binds to the TLR domain of MYD88. DHX36 is a component of a cytosolic viral sensor and is recruited to a complex consisting of DDX1, DDX21 and TICAM1, which triggers type I interferon and
62011	DHX36	21703541	<ul><li>cytokine response to dsRNA. (Demonstrated in murine model)</li><li>Ddx21 is a component of a cytosolic viral sensor and is recruited to a complex consisting of Ddx1, Dhx36 and</li></ul>
157719	Ddx21	21703541	Ticam1, which triggers type I interferon and cytokine response to dsRNA. DDX21 is a component of a cytosolic viral sensor and is recruited to a complex consisting of DDX1, DHX36 and TICAM1, which triggers type I interferon and
76283	DDX21	21703541	cytokine response to dsRNA. (Demonstrated in murine model)
			Ddx1 is a cytosolic viral sensor in dendritic cells that binds to dsRNA through its Helicase A domain. Ddx1 then recruits Ddx21, Dhx36 and Ticam1 to mount type I interferon and cytokine response to poly I:C,
128974	Ddx1	21703541	influenza A virus, and reovirus. DDX1 is cytosolic viral sensor in dendritic cells that binds to dsRNA through its Helicase A domain. DDX1 then recruits DDX21, DHX36 and TICAM1 to mount type I interferon and cytokine response to poly I:C,
30761	DDX1	21703541	Influenza A virus and reovirus. (Demonstrated in murine model)
172600	Cd97	21706400	Cd97 plays an important role in recruiting granulocytes and possibly macrophages to the sites of infection. CD97 plays an important role in recruiting
33120	CD97	21706400	granulocytes and possibly macrophages to the sites of infection. Activation of Adrb2 on Nod2/Tlr2-stimulated dendritic
152615	Adrb2	21683614	cells biases the cell priming ability towards an Th17 immune response. Activation of ADRB2 on NOD2/TLR2-stimulated dendritic cells biases the cell priming ability towards an Th17 immune response. (Demonstrated in murine
52685	ADRB2	21683614	model)
145910	Jam3	21706006	Jam3 is a key regulator of polarized neutrophil transendothelial migration in vivo. JAM3 is a key regulator of polarized neutrophil
76413	JAM3	21706006	transendothelial migration in vivo. (Demonstrated in murine model) AHR deficiency imapirs TLR and NFkB-mediated proinflammatory gene expression after activation by a classical stimulus, such as LPS. (Demonstrated in
8995	AHR	21683686	murine model)

			Hifla, under normoxic conditions, accumulates in dendritic cells via the TLR/Myd88/NFkB signalling pathway to induce a distinct subset of proinflammatory
148620	Hifla	21685248	genes in comparison to hypoxia-induced Hifla. Ubqln1 strongly suppresses the transcriptional activation of Ifnb promoter and is an inhibitor of the
157923	Ubqln1	21695056	Tlr3/Ticam1 anti-viral pathway by reducing Ticam1 protein levels. (Demonstrated in human) UBQLN1 strongly suppresses the transcriptional activation of the IFNB promoter and is an inhibitor of
73620	UBQLN1	21695056	the TLR3/TICAM1 anti-viral pathway by reducing TICAM1 protein levels. Pmaip1 is induced by the RIG-I/MDA5 axis during
155006	Pmaip1	21698224	viral infection and activates apoptosis in macrophages, dendritic cells and primary lymphocytes. PMAIP1 is induced by the RIG-I/MDA5 axis during viral infection and activates apoptosis in macrophages,
4142	PMAIP1	21698224	dendritic cells and primary lymphocytes. Ip6k1 disruption results in the augmentation of downstream phosphatidylinositol-(3,4,5)-triphosphate signalling in neutrophils. As a result, these neutrophils exhibited greater phagocytic and bactericidal ability and amplified NADPH oxidase-mediated production of
198951	Ip6k1	21685907	superoxide. IP6K1 disruption results in the augmentation of downstream phosphatidylinositol-(3,4,5)-triphosphate signalling in neutrophils. As a result, these neutrophils exhibited greater phagocytic and bactericidal ability
35427	IP6K1	21685907	<ul> <li>and amplified NADPH oxidase-mediated production of superoxide.</li> <li>Khsrp directly interacts with AU-rich elements in the 3' UTR of Ifn mRNA to negatively regulate the transcript levels. Khsrp deficient mouse embryonic fibroblast produced higher levels of Ifna4 and Ifnb mRNAs in response to viral infections as a result of decreased</li> </ul>
193424	Khsrp	21690298	mRNA decay. KHSRP directly interacts with AU-rich elements in the 3' UTR of IFN mRNA to negatively regulate the transcript levels. Khsrp deficient mouse embryonic fibroblast produced higher levels of Ifna4 and Ifnb
21386	KHSRP	21690298	mRNAs in response to viral infections as a result of decreased mRNA decay. Ifnar1 deficiency completely abolishes the reduction of sterol biosynthetic activity of macrophages during viral infections, thereby linking the regulation of lipid metabolism pathway with interferon anti-viral defence
176038	Ifnar1	21408089	responses. Ifnar1 is phosphorylated by p38 MAP kinase in response to pathogen-recognition receptor stimulation. This phosphorylation promotes Ifnar1 ubiquitination and accelerates the proteolytic turnover of the receptor,
176038	Ifnar1	21695243	which leads to attenuation of type I IFN signalling.
			IFNAR1 acts as a docking site for the latent form of STAT2 and mediates the interaction between JAK
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2095	IFNAR1	8605876	kinases and the STAT transcription factors. IFNAR1 interacts with the amino-terminal half of
2095	IFNAR1	9461596	TYK2 and this interaction is required for interferon (IFN)-alpha signal transduction. The interferon alpha receptor is composed of two
			subunits: IFNAR1 and IFNAR2;IL10RB. IFNAR1
2095	IFNAR1	12220192	binds to STAT2 to initiate interferon (IFN) signalling. Type I interferons (IFNs) play an important role in innate immunity to protozoan parasites by binding the IFN alpha receptor, composed of IFNAR1 and IFNAR2;IL10RB, and regulating neutrophil/monocyte recruitment, neutrophil turnover, and Leishmania
2095	IFNAR1	20483775	infection. IFNAR1 deficiency completely abolishes the reduction
			of sterol biosynthetic activity of macrophages during viral infections, thereby linking the regulation of lipid
2095	IFNAR1	21408089	metabolism pathway with interferon anti-viral defence responses. (Demonstrated in murine model) IFNAR1 is phosphorylated by p38 MAP kinase in
			response to pathogen-recognition receptor stimulation. This phosphorylation promotes IFNAR1 ubiquitination
2095	IFNAR1	21695243	and accelerates the proteolytic turnover of the receptor, which leads to attenuation of type I IFN signalling.
202215	Tpst1	21665277	Tpst1 silencing reduces the II6 production in LPS- stimulated macrophages. TPST1 silencing reduces the IL6 production in LPS-
18615	TPST1	21665277	stimulated macrophages. (Demonstrated in murine model)
			Plec silencing reduces the Il6 production in LPS-
153069	Plec	21665277	stimulated macrophages. PLEC silencing reduces the IL6 production in LPS-
39382	PLEC	21665277	stimulated macrophages. (Demonstrated in murine model)
			VENTX plays a pivotal role in human macrophage terminal differentiation and proinflammatory function. VENTX is upregulated during monocyte to macrophage differentiation, and ablation of VENTX in monocytes profoundly impairs their differentiation to
93605	VENTX	21670496	macrophages. Epithelial Ccr3 is involved in the progression of LPS-
206550	Ccr3	21660963	induced lung inflammation by mediating the release of Il8. Epithelial CCR3 is involved in the progression of LPS-
30125	CCR3	21660963	induced lung inflammation by mediating the release of IL8. Vldlr is exploited by rhinovirus for viral entry. Vldlr
155959	Vldlr	21642441	protein levels are negatively regulated by IFN/RIG-I signalling via MIR23B. (Demonstrated in human)

44752	VLDLR	21642441	<ul><li>VLDLR is exploited by rhinovirus for viral entry.</li><li>VLDLR protein levels are negatively regulated by IFN/RIG-I signalling via MIR23B.</li><li>MIR23B expression is induced by RIG-I signalling and targets VLDLR and LRP5. MIR23B specifically</li></ul>
127077	MIR23B	21642441	inhibits rhinovirus infection which utilizes VLDLR for viral entry. Cflar confers protection against cytoplasmic dsRNA-
157242	Cflar	21(25782	mediated cell death and down-regulates Irf3- and NFkB-mediated gene expression. In addition, Cflar also negatively regulates LPS-induced Tlr4-signalling in endothelial cells and protects against Tlr4-mediated
157342	Cliar	21635783	apoptosis. CFLAR confers protection against cytoplasmic dsRNA-mediated cell death and down-regulates IRF3- and NFkB-mediated gene expression. In addition,
			CFLAR also negatively regulates LPS-induced TLR4- signalling in endothelial cells and protects against
78386	CFLAR	21635783	TLR4-mediated apoptosis. (Demonstrated in murine model)
			Havcr2 is constitutively expressed on human resting monocytes/macrophages and functions as a cap to block Il12, which is a key pro-inflammatory cytokine linking innate and adaptive immune responses. Havcr2 plays a crucial role in the negative regulation of innate immune responses through crosstalk with Pdcd1 and
165782	Haver2	21637332	Socs1 to limit Stat1 phosphorylation in HCV infection. (Demonstrated in human)
105782	Haverz	21057552	Cdk6 expression is induced upon LPS stimulation via Tlr4-signalling pathway. Cdk6 mediates the increased adhesion of macrophages in response to LPS challenge,
131439	Cdk6	21628465	and is important for LPS lethality. CDK6 expression is induced upon LPS stimulation via TLR4-signalling pathway. CDK6 mediates the increased adhesion of macrophages in response to LPS challenge, and is important for LPS lethality.
26773	CDK6	21628465	(Demonstrated in murine model) MIR107 is a negative regulator of macrophage adhesion by targeting the degradation of CDK6 RNA, which confers protection against the lethal effect of
126455	MIR107	21628465	LPS. The expression of MIR107 is inhibited by TLR4 signalling. Gpr77 is an alternate receptor for C5a and is negatively regulated by the TLR-signalling pathway. TLR-mediated inhibition of Gpr77 results in the
146283	Gpr77	21630250	amplification of complement pro-inflammatory responses. GPR77 is an alternate receptor for C5a and is negatively regulated by the TLR-signalling pathway. TLR-mediated inhibition of GPR77 results in the
59470	GPR77	21630250	amplification of complement pro-inflammatory responses. (Demonstrated in murine model)

			Lilrb3 negatively regulates macrophage functions in response to pathogenic bacteria and chronic intestinal inflammatory responses, as demonstrated by increased production of proinflammatory cytokines (IL-6, IL- 1beta and TNF-alpha) and activation of MAPK and
136357	Lilrb3	20398663	NFkappaB in Lilrb3(-/-) macrophages following bacterial activation. Pik3ap1 acts as a negative regulator of TLR-induced production of IL6 and IL10 in macrophages in response
165835	Pik3ap1	20728433	to LPS stimulation. PIK3AP1 acts as a negative regulator of TLR-induced
83923	PIK3AP1	20728433	production of IL6 and IL10 in macrophages in response to LPS stimulation. (Demonstrated in murine model) Avp regulates water absorption in the collecting duct and acts as a potent modulator of the TLR4-mediated intrarenal innate response caused by uropathogenic E.
206576	Avp	21615666	coli. AVP regulates water absorption in the collecting duct and acts as a potent modulator of the TLR4-mediated
45432	AVP	21615666	<ul><li>intrarenal innate response caused by uropathogenic E.</li><li>coli.</li><li>Casp8 restricts the RIG-I-mediated activation of Irf3 by catalytically cleaving Ripk1, and subsequently</li></ul>
157464	Casp8	21419663	converting Ripk1 from a signalling enhancer to a signalling inhibitor. CASP8 has an essential role in the regulation of NF- kappaB function in response to TLR4 stimulation, whereby CASP8 recruitment to IKBKB leads to
78534	CASP8	17213198	delayed NF-kappaB nuclear translocation and impaired NF-kappaB transcriptional activity.
78534	CASP8	20019748	CASP8 is involved in pro-apoptotic signalling through TLR3 and its activation is TICAM1 (TRIF)-dependent. CASP8 restricts the RIG-I-mediated activation of IRF3 by catalytically cleaving RIPK1, and subsequently converting RIPK1 from a signalling enhancer to a
78534	CASP8	21419663	signalling inhibitor. SREBF1 deficient mice are resistant to endotoxic shock and systemic inflammatory response syndrome induced by cecal ligation and puncture. SREBF1 is not only a necessary transcription factor for lipogenesis genes in macrophages, but is also responsible for the
33208	SREBF1	21531336	expression of NLRP1A, which is a core inflammasome component. (Demonstrated in murine model) Srebf1 deficient mice are resistant to endotoxic shock and systemic inflammatory response syndrome induced by cecal ligation and puncture. Srebf1 is not only necessary for transcription of lipogenesis genes in macrophages but is also responsible for the expression
179865	Srebfl	21531336	macrophages, but is also responsible for the expression of Nlrp1a, which is a core inflammasome component.

			Zc3hav1 isoform ZAPS is selectively induced by 5'- triphosphate-modified RNA and functions as a potent
137011	Zc3hav1	21102435	stimulator of interferon responses via the RIG-I signalling pathway. ZC3HAV1 isoform ZAPS is selectively induced by 5'-
43487	ZC3HAV1	21102435	triphosphate-modified RNA and functions as a potent stimulator of interferon responses via the RIG-I signalling pathway.
			ATG5 forms a conjugate with ATG12 to directly associate with DDX58 and MAVS and negatively regulate the antiviral IFN production pathway by
94856	ATG5	17709747	mediating autophagy. ATG12::ATG5 conjugate is a key regulator of the
			autophagic process to eliminate pathogens such as Streptococcus, M. tuberculosis, Listeria, and herpesvirus. ATG12::ATG5 also associates with components of the RIG-I pathway to negatively regulate type I IFN response and promote RNA virus
94856	ATG5	17921696	replication. Atg12::Atg5 conjugate is a key regulator of the autophagic process to eliminate pathogens such as Streptococcus, M. tuberculosis, Listeria, and herpesvirus. Atg12::Atg5 also associates with components of the RIG-I pathway to negatively
145432	Atg12	17921696	regulate type I IFN response and promote RNA virus replication. ATG12::ATG5 conjugate is a key regulator of the autophagic process to eliminate pathogens such as Streptococcus, M. tuberculosis, Listeria, and herpesvirus. ATG12::ATG5 also associates with
			components of the RIG-I pathway to negatively regulate type I IFN response and promote RNA virus
37900	ATG12	17921696	replication. MIR373 is upregulated in hepatitis B virus-infected
126503	MIR373	21608007	liver tissues to target NFIB mRNA for degradation and promotes viral replication. MIR372 is upregulated in hepatitis B virus-infected
127071	MIR372	21608007	liver tissues to target NFIB mRNA for degradation and promotes viral replication. Akna targeted deletion results in susceptibility to pathogen-induced systemic inflammation and causes
154783	Akna	21606955	sudden neonatal death.
			AKNA targeted deletion results in susceptibility to pathogen-induced systemic inflammation and causes sudden neonatal death. (Demonstrated in murine model)
82103	AKNA	21606955	IRAK2 has a role in TRAF6 ubiquitination in the Toll- like receptor (TLR) pathway and thus plays a more
17914	IRAK2	17878161	central role than IRAK1 in TLR signalling to NFkappaB.

17014		10420411	IRAK2 is critical in late-phase Toll-like receptor (TLR) responses, and IRAK1 and IRAK2 are essential for the initial responses to TLP atimulation
17914	IRAK2	18438411	initial responses to TLR stimulation. IRAK2 is required for LPS-mediated post- transcriptional control of cytokine and chemokine expression, which plays an essential role in TLR4-
17914	IRAK2	19224918	induced septic shock. IRAK2 induces cytokine and chemokine mRNA stability and translation in response to LPS stimulation in macrophages. The kinase activity of IRAK2 is required for the optimal activation of mitogen activated protein kinase signaling, which regulates cytokine/chemokine production at posttranscriptional levels.
17914	IRAK2	21291324	
			IRAK2 is required for both TLR4 and TLR8-mediated activation of NFkB and p38 MAP kinase, and the induction of TNF mRNA. In addition, IRAK2 is required for regulating MYD88-dependent TNF-alpha mRNA stability. (Phenotype was not observed in
17914	IRAK2	21606490	murine ortholog, Irak2) The JAK2/STAT5/CIS pathway suppresses CCL1 but
40784	CCL1	18981157	not CCL7 and CCL8 chemokine expression CCL1 secretion in freshly isolated monocytes is induced by the combined engagement of TLR3/TLR4/TLR8. As monocytes differentiate, the capacity to induce CCL1 is lost via a IL23-dependent mechanism.
40784	CCL1	21601943	
			Pglyrp4 is a member of the Peptidoglycan Recognition Proteins (PGRP) family and recognizes peptidoglycan, a structural component of bacterial cell walls, as a part
168143	Pglyrp4	21439073	of innate immune response against infections. Pglyrp4 binds to Gram-positive bacterial wall and activates a protein-sensing two-component system to induce bacterial death. PLGYRP1-mediated activation results in membrane depolarization and cessation of peptidoglycan, protein, and RNA/DNA synthesis, as well as the production of hydroxyl radicals. (Demonstrated in human)
168143	Pglyrp4	21602801	
			PGLYRP4 is a secreted innate immunity protein that is expressed on body surfaces, mucous membranes, and in secretions (saliva, sweat) and is conserved from insects to mammals, it recognizes bacterial peptidoglycan, and functions in antibacterial immunity and inflammation.
102639	PGLYRP4	20418257	

102639	PGLYRP4	21439073	PGLYRP4 is a member of the Peptidoglycan Recognition Proteins (PGRP) family and recognizes peptidoglycan, a structural component of bacterial cell walls, as a part of innate immune response against infections. PGLYRP4 binds to Gram-positive bacterial wall and activates a protein-sensing two-component system to induce bacterial death. PLGYRP1-mediated activation results in membrane depolarization and cessation of peptidoglycan, protein, and RNA/DNA synthesis, as well as the production of hydroxyl radicals.
102639	PGLYRP4	21602801	
148760	Dalumi	21134971	Pglyrp1 is a peptidoglycan recognition protein and play a role in innate immunity against L. monocytogenes infection by inducing Tnfa.
148/00	Pglyrp1	211349/1	Pglyrp1 is a member of the Peptidoglycan Recognition
148760	Pglyrp1	21439073	<ul> <li>Proteins (PGRP) family and recognizes peptidoglycan, a structural component of bacterial cell walls, as a part of innate immune response against infections.</li> <li>Pglyrp1 binds to Gram-positive bacterial wall and activates a protein-sensing two-component system to induce bacterial death. PLGYRP1-mediated activation results in membrane depolarization and cessation of peptidoglycan, protein, and RNA/DNA synthesis, as well as the production of hydroxyl radicals. (Demonstrated in human)</li> </ul>
148760	Pglyrp1	21602801	(Demonstrated in numan)
			Gbp10 is an IFN gamma-inducible gene that confers cell-autonomous immunity to listerial or mycobacterial infection within macrophages. Gbp10 interacts with host defense proteins, including phagocyte oxidase, antimicrobial peptides and autophagy effectors, to kill
185854	Gbp10	21551061	intracellular bacteria. Gbp7 is an IFN gamma-inducible gene that confers cell-autonomous immunity to listerial or mycobacterial infection within macrophages. Gbp7 interacts with host defense proteins, including phagocyte oxidase, antimicrobial peptides and autophagy effectors, to kill intracellular bacteria.
197215	Gbp7	21551061	Gbp6 is an IFN gamma-inducible gene that confers cell-autonomous immunity to listerial or mycobacterial infection within macrophages. Gbp6 interacts with host defense proteins, including phagocyte oxidase,
255448	Gbp6	21551061	antimicrobial peptides and autophagy effectors, to kill intracellular bacteria.

			Gbp1 is an IFN gamma-inducible gene that confers cell-autonomous immunity to listerial or mycobacterial infection within macrophages. Gbp1 interacts with host defense proteins, including phagocyte oxidase, antimicrobial peptides and autophagy effectors, to kill intracellular bacteria.
197289	Gbp1	21551061	Tlr2 and Tnfsfl1 signalling pathways are modulated by Porphromonas gingivalis to alter the differentiation states of osteoclasts resulting in bacteria-mediated bone loss.
182101	Tnfsfl 1	21566133	TLR2 and TNFSF11 signalling pathways are modulated by Porphromonas gingivalis to alter the differentiation states of osteoclasts resulting in bacteria-mediated bone loss. (Demonstrated in murine model)
28393	TNFSF11	21566133	Exogenous IL4 was sufficient to drive the accumulation of tissue macrophages through self-renewal revealing that the expansion of innate cells necessary for pathogen control or wound repair can occur without recruitment of potentially tissue-destructive inflammatory cells. (Demonstrated in murine model)
42701	IL4	21566158	Arhgap15 negatively modulates Akt1 activity and thereby negatively regulates neutrophil function. Arhgap15 deficiency results in increased neutrophil recruitment to the site of infection and offers protection against an experimental model of severe abdominal sepsis.
168042	Arhgap15	21551229	ARHGAP15 negatively modulates AKT1 activity and thereby negatively regulates neutrophil function. ARHGAP15 deficiency results in increased neutrophil recruitment to the site of infection and offers protection against an experimental model of severe abdominal sepsis. (Demonstrated in murine model)
71605	ARHGAP15	21551229	Bid is a critical component of the NOD signalling pathway and is crucial for the peptidoglycan inflammation response. Bid deficient mice are unresponsive to local or systemic exposure to NOD agonist or their protective effects in experimental colitis.
184076	Bid	21552281	contris.

			BID is a critical component of the NOD signalling pathway and is crucial for the peptidoglycan inflammation response. BID deficient mice are unresponsive to local or systemic exposure to NOD agonist or their protective effects in experimental colitis. (Demonstrated in murine model)
1050	BID	21552281	Thbs1 expression is associated with decreased phagocytosis and bacterial clearance, resulting in increased peritoneal inflammation and mortality from sepsis.
197807	Thbs1	21573017	THBS1 expression is associated with decreased phagocytosis and bacterial clearance, resulting in increased peritoneal inflammation and mortality from sepsis. (Demonstrated in murine model)
5926	THBS1	21573017	Tecpr1 activity is necessary for efficient autophagic targeting of bacteria and the suppression of Shigella intracellular replication.
208049	Tecpr1	21575909	TECPR1 activity is necessary for efficient autophagic targeting of bacteria and the suppression of Shigella intracellular replication. (Demonstrated in murine model)
28493	TECPR1	21575909	HMGN2 acts as a positive modulator of NF-kB signalling to promote LPS-induced beta-defensin expression. HMGN2 prolongs the retention time and enhances the accumulation of NF-kB p65 and synergistically bind to the DEFB4A (HBD-2)
94677	HMGN2	21518253	promoter to induce expression. CXCR3 expression on recruited peritoneal macrophages and granulocytes increases following sepsis, and deletion of CXCR3 significantly increases mortality to a septic challenge in neonatal mice.
76333	CXCR3	21518789	(Demonstrated in murine model) Map3k5 is required for LPS-induced activation of p38, which is a crucial determinant of the production of pro- inflammatory cytokines in endotoxemia.
136255	Map3k5	21515258	MAP3K5 is required for LPS-induced activation of p38, which is a crucial determinant of the production of pro-inflammatory cytokines in endotoxemia. (Demonstrated in murine model)
96934	MAP3K5	21515258	PTGES (prostaglandin E2) is produced by LPS-primed macrophages upon treatment with silica crystal and aluminum salts, and is important for the production of
89226	PTGES	21497116	IgE in Th2 cells. (Demonstrated in murine model)

			Grn is an essential secreted cofactor that potentiates Tlr9-driven response to CpG olignonucletides by binding directly to CpG oligos and Tlr9. Grn deficient
212642	Grn	21497117	<ul><li>murine macrophages showed impaired delivery of CpG</li><li>oligos to endolysosomal compartments and reduced</li><li>response to CpG.</li><li>GRN is an essential secreted cofactor that potentiates</li></ul>
			TLR9-driven response to CpG olignonucletides by binding directly to CpG oligos and TLR9. GRN deficient murine macrophages showed impaired delivery of CpG oligos to endolysosomal compartments and reduced response to CpG.
53782	GRN	21497117	(Demonstrated in murine model) Tlr2::Tlr6 synergistically interacts with Tlr9 in lung epithelium to induce rapid pathogen killing, and can be used as a therapeutic target to treat otherwise lethal
165921	Tlr6	21482737	pneumonia. Cd46 enhances nitric oxide production in mouse
209330	Cd46	10233938	<ul><li>macrophages in response to measles virus infection in the presence of gamma interferon.</li><li>Cd46 is expressed ubiquitously and functions as a co-factor in the factor I-mediated proteolytic cleavage of C3b and C4b. Cd46 has a vital role in preventing</li></ul>
209330	Cd46	21488871	complement deposition on host tissue, and the resulting auto-immunity. CD46 is a cell surface pathogen receptor that induces autophagy upon pathogen recognition and this CD46-
106372	CD46	20087059	<ul><li>dependent autophagy is critical for early control of the infection.</li><li>CD46 is a ubiquitously expressed membrane protein that regulates complement activation, as a cellular attachment receptor of several pathogens, including</li></ul>
106372	CD46	15919905	<ul><li>measles virus, Neisseria gonorrhea, adenovirus and human herpesvirus 6.</li><li>CD46 enhances bacterial survival and represents a</li></ul>
106372	CD46	18573902	novel pathogenic mechanism that contributes to the severity of group A streptococcal disease. CD46 acts as a human epithelial cell receptor for
106372	CD46	16888016	<ul><li>internalization of opsonized uropathogenic Escherichia coli.</li><li>CD46 plays a key role in tailoring innate immune</li></ul>
			recognition of apoptotic and necrotic cells, whereby the complement innate immune system is using two synergistic strategies with the recognition of altered self-nucleic acids (NA) and missing self-CD46 signals
106372	CD46	16087667	to instruct and tailor the efficient removal of apoptotic and necrotic cells in immunoprivileged sites. CD46 is expressed ubiquitously and functions as a co- factor in the factor I-mediated proteolytic cleavage of C3b and C4b. CD46 has a vital role in preventing
106372	CD46	21488871	complement deposition on host tissue, and the resulting auto-immunity.

FCN1, as well as ficolins FCN2 and FCN3, in serum are associated with MBL-associated serine protease (MASP) to form a complex and this complex binds to carbohydrates present on the surface of a variety of Gram-positive and Gram-negative bacteria through ficolin, initiating complement activation via the lectin pathway.

91827	FCN1	20375620	pathway.
91827	FCN1	16116205	FCN1, like its family members, functions as a recognition molecule of the lectin complement pathway and plays an important role in innate immunity. FCN3 and PTX3 are soluble oligomeric pattern-recognition molecules that interact with each other and
91827	FCN1	21490156	act synergistically to activate the lectin complement pathway. Nr1h3 repressed Irf3- or Irf7-induced transactivation of
190792	Nr1h3	21492741	the interferon-beta promoter and NDV infection further potentiated the repressive effect in dendritic cells. NR1H3 repressed IRF3- or IRF7-induced transactivation of the interferon-beta promoter and
42877	NR1H3	21492741	NDV infection further potentiated the repressive effect in dendritic cells. (Demonstrated in murine model) Nr4a3 repressed Irf3- or Irf7-induced transactivation of
146207	Nr4a3	21492741	the interferon-beta promoter and NDV infection further potentiated the repressive effect in dendritic cells. NR4A3 repressed IRF3- or IRF7-induced transactivation of the interferon-beta promoter and
78601	NR4A3	21492741	NDV infection further potentiated the repressive effect in dendritic cells. (Demonstrated in murine model) H2-Ab1 is an intracellular MHC class II molecule that
170861	H2-Ab1	21441935	can act as an adaptor to promote the full activation of the TLR-triggered innate immune response. H2-Aa is an intracellular MHC class II molecule that
170982	H2-Aa	21441935	can act as an adaptor to promote the full activation of the TLR-triggered innate immune response. Srxn1 is transcriptionally regulated by Nfe2l2 in immunostimulated primary macrophages that produce both reactive oxygen species and nitric oxide. The nitric oxide/Nfe2l2/Srxn1 pathway participates in the maintenance of redox homeostasis in cytokine-
209605	Srxn1	21466852	activated macrophages and other inflammatory settings. SRXN1 is transcriptionally regulated by NFE2L2 in immunostimulated primary macrophages that produce both reactive oxygen species and nitric oxide. The nitric oxide/NFE2L2/SRXN1 pathway participates in the maintenance of redox homeostasis in cytokine- activated macrophages and other inflammatory
38297	SRXN1	21466852	settings. (Demonstrated in murine model)

			Gp2 on M cells (specialized epithelial antigen- transporting cells) functions as an uptake receptor for a subset of commensal and pathogenic bacteria. Gp2 interacts with type 1 pilus of gram-negative
207909	Gp2	21468225	enterobacilli such as E. coli and Salmonella enterica. GP2 on M cells (specialized epithelial antigen- transporting cells) functions as an uptake receptor for a subset of commensal and pathogenic bacteria. GP2 interacts with type 1 pilus of gram-negative enterobacilli such as E. coli and Salmonella enterica.
18165	GP2	21468225	<ul> <li>(Demonstrated in murine model)</li> <li>Ripk2 is a downstream adaptor molecule in the Nod1/2 signalling pathway and is important for the progression and pathogenesis of experimental autoimmune encephalomyelitis (animal model of multiple sclerosis).</li> <li>Ripk2 was found to be critical for the activation of CNS-infiltrating dendritic cells.</li> </ul>
130745	Ripk2	21236705	Ripk2 is essential for Nod1 and Nod2-signalling upon recognition of bacterial peptidoglycan. Ripk2 is crucial for inflammatory cytokine secretion, activation and
130745	Ripk2	21469090	recruitment of macrophage and neutrophils as well as the capacity to activate the adaptive immune response. Hrh4 is a histamine receptor that can mediate cytokine production (e.g. Il6) from mast cells. Hrh4 is also able to synergize with other inflammatory signals, such as
129289	Hrh4	21469095	LPS, to potentiate cytokine production and contribute to inflammation. HRH4 is a histamine receptor that can mediate cytokine production (e.g. IL6) from mast cells. HRH4 is also able to synergize with other inflammatory signals, such as LPS, to potentiate cytokine production and contribute to inflammation. (Demonstrated in
1754	HRH4	21469095	murine model) Plg is a serum protein that interacts with B. anthracis spores and cleaves complement 3 molecules, resulting in a decrease in macrophage phagocytosis.
136615	Plg	21464960	(Demonstrated in human) PLG is a serum protein that interacts with B. anthracis spores and cleaves complement 3 molecules, resulting
98749	PLG	21464960	in a decrease in macrophage phagocytosis. Ccl2 is secreted by bone marrow mesenchymal stem cells in response to circulating TLR ligands or bacterial infection, which then induces monocyte trafficking into
204473	Ccl2	21458307	the bloodstream. CCL2 is secreted by bone marrow mesenchymal stem cells in response to circulating TLR ligands or bacterial infection, which then induces monocyte trafficking into
40504	CCL2	21458307	the bloodstream. (Demonstrated in murine model)

			Impdh2 proteins are rapidly recruited to the lipid raft of monocytes after lipopeptide stimulation, and play an essential role in the negative regulation of Tlr2 signalling by modulating PI3K activity. (Demonstrated
200174	Impdh2	21460227	in human) IMPDH2 proteins are rapidly recruited to the lipid raft of monocytes after lipopeptide stimulation, and play an essential role in the negative regulation of TLR2
34157	IMPDH2	21460227	signalling by modulating PI3K activity. C1qa :: C1qb :: C1qc (C1q) is a versatile innate immune molecule that recognizes an array of self, non- self and altered-self ligands. The broad-spectrum of ligand specificity is facilitated by the modular organization of the heterotrimeric globular region and
198256	Clqc	21450789	its ability to change its confirmation. C1QC is the C-chain of the C1Q recognition subunit of Complement component 1 (C1), and acts a multimolecular protease that triggers the classical pathway of complement and has a major role in the
93737	C1QC	15207504	host defence against pathogens. C1Q is involved in the modulation of various immune cells such as dendritic cells, platelets, microglia cells and lymphocytes. C1Q has roles in clearance of apoptotic cells as well as a range of cell processes such as differentiation, chemotaxis, aggregation and adhesion, and pathogenesis of neurodegenerative
93737	C1QC	20381531	diseases. C1QA :: C1QB :: C1QC (C1Q) is a versatile innate immune molecule that recognizes an array of self, non- self and altered-self ligands. The broad-spectrum of ligand specificity is facilitated by the modular
93737	C1QC	21450789	organization of the heterotrimeric globular region and its ability to change its confirmation. Clqa :: Clqb :: Clqc (Clq) is a versatile innate immune molecule that recognizes an array of self, non- self and altered-self ligands. The broad-spectrum of ligand specificity is facilitated by the modular organization of the heterotrimeric globular region and
198224	Clqb	21450789	its ability to change its confirmation. C1QB is the B-chain of the C1Q, the recognition subunit of Complement component 1 (C1), is a multimolecular protease that triggers the classical pathway of complement and has a major role in the
93747	C1QB	15207504	host defence against pathogens. C1Q is involved in the modulation of various immune cells such as dendritic cells, platelets, microglia cells and lymphocytes. C1Q has roles in clearance of apoptotic cells as well as a range of cell processes such as differentiation, chemotaxis, aggregation and
93747	C1QB	20381531	adhesion, and pathogenesis of neurodegenerative diseases.

93747	C1QB	21450789	C1QA ::: C1QB ::: C1QC (C1Q) is a versatile innate immune molecule that recognizes an array of self, non- self and altered-self ligands. The broad-spectrum of ligand specificity is facilitated by the modular organization of the heterotrimeric globular region and its ability to change its confirmation. C1qa :: C1qb :: C1qc (C1q) is a versatile innate immune molecule that recognizes an array of self, non- self and altered-self ligands. The broad-spectrum of
198277	C1qa	21450789	ligand specificity is facilitated by the modular organization of the heterotrimeric globular region and its ability to change its confirmation.
198277	Ciqa	21430789	C1QA is the A-chain of the C1Q recognition subunit of Complement component 1 (C1), and acts a multimolecular protease that triggers the classical pathway of complement and has a major role in the
93718	C1QA	15207504	host defence against pathogens. C1Q is involved in the modulation of various immune cells such as dendritic cells, platelets, microglia cells
			and lymphocytes. C1Q has roles in clearance of apoptotic cells as well as a range of cell processes such as differentiation, chemotaxis, aggregation and adhesion, and pathogenesis of neurodegenerative
93718	C1QA	20381531	diseases. C1QA :: C1QB :: C1QC (C1Q) is a versatile innate immune molecule that recognizes an array of self, non- self and altered-self ligands. The broad-spectrum of ligand specificity is facilitated by the modular
93718	C1QA	21450789	organization of the heterotrimeric globular region and its ability to change its confirmation. Ccbp2 is a marker for innate-like B cells, which are a
205741	Ccbp2	21450903	heterogeneous collection of cells that control infection and suppress inflammation. Ccbp2 act as a chemokine scavenger receptor by internalizing chemokines without inducing calcium fluxes or chemotaxis in innate B cells.
			CCBP2 is a marker for innate-like B cells, which are a heterogeneous collection of cells that control infection and suppress inflammation. CCBP2 act as a chemokine scavenger receptor by internalizing chemokines
28131	CCBP2	21450903	without inducing calcium fluxes or chemotaxis in innate B cells. (Demonstrated in murine model) Nfkbia degradation occurs through the TNF-stimulated
140708	Nfkbia	21454695	formation of autophagosomes in epithelial cells, which results in the prolonged activation of NFKB activity. Xiap facilitates ubiquitin-dependent signalling activated by pattern recognition receptors, such as TLR and NOD, to mediate the activation of NFKB
141324	Xiap	21447281	transcription.
85142	XIAP	18769721	XIAP regulates cytosol-specific innate immunity to Listeria infection.

85142	XIAP	19667203	XIAP interacts with NOD1 and NOD2 and mediates NOD signalling via interaction with RIPK2. XIAP and its E3 ligase activity promote transforming
85142	XIAP	19531477	growth factor-{beta}-mediated NF-kappaB activation during breast cancer progression. X-linked inhibitor of apoptosis protein (XIAP) in a
85142	XIAP	20406824	complex with survivin, a physiological substrate for granzyme M (GzmM), act to inhibit caspase activation. XIAP facilitates ubiquitin-dependent signalling activated by pattern recognition receptors, such as TLR
85142	XIAP	21447281	and NOD, to mediate the activation of NFKB transcription. Birc3 facilitates ubiquitin-dependent signalling
133177	Birc3	21447281	activated by pattern recognition receptors, such as TLR and NOD, to mediate the activation of NFKB transcription. Birc2 facilitates ubiquitin-dependent signalling
133125	Birc2	21447281	activated by pattern recognition receptors, such as TLR and NOD, to mediate the activation of NFKB transcription. BIRC2 regulates TNF-alpha(TNF)-mediated NF-
69075	BIRC2	18697935	kappaB activation by binding to TNFRSF1A (TNF receptor 1). BIRC2 is the ubiquitin protein ligase for ASK1
			ubiquitination, a protein that plays an essential role in tumour necrosis factor alpha (TNF-alpha)-induced mitogen-activated protein kinase signalling. BIRC2 is
69075	BIRC2	17220297	also responsible for regulating the duration of TNF signalling in primary cells expressing TNFR2. BIRC2 and BIRC3 are required for innate immunity
69075	BIRC2	19464198	signalling by the pattern recognition receptors NOD1 and NOD2. BIRC2 plays a role in lipopolysaccharide (LPS)-
69075	BIRC2	20458734	induced autophagy in vascular endothelial cells (VECs). BIRC2 facilitates ubiquitin-dependent signalling
69075	BIRC2	21447281	activated by pattern recognition receptors, such as TLR and NOD, to mediate the activation of NFKB transcription. Hspala is secreted into the extracellular space during exercise-induced stress and increases the intracellular
175332	Hspala	21448922	levels of cAMP, which acts as an "intracellular danger signal" to activate neutrophils. (Demonstrated in human) Pglyrp2 is a member of the Peptidoglycan Recognition Proteins (PGRP) family and recognizes peptidoglycan,
251732	Pglyrp2	21439073	a structural component of bacterial cell walls, as a part of innate immune response against infections. PGLYRP2 is a secreted innate immunity protein that is expressed in the liver and is conserved from insects to
34754	PGLYRP2	20418257	mammals, it recognizes bacterial peptidoglycan, and functions in antibacterial immunity and inflammation.

			PGLYRP2 is a member of the Peptidoglycan Recognition Proteins (PGRP) family and recognizes peptidoglycan, a structural component of bacterial cell walls, as a part of innate immune response against
34754	PGLYRP2	21439073	infections. Defb3 expression is inhibited in keratinocytes under high glucose conditions, which in turn contributed to the frequent occurrences of infection associated with
137962	Defb3	21442129	<ul><li>diabetic wounds. (Demonstrated in rat model and in human)</li><li>Raet1c expression on erythroblast surface is induced early after Friend virus inoculation, and is recognized by Klrk1 expressed on the NK cells to trigger cytotoxic</li></ul>
137529	Raet1c	21411527	activities. Raet1a expression on erythroblast surface is induced early after Friend virus inoculation, and is recognized
270707	Raet1a	21411527	<ul><li>by Klrk1 expressed on the NK cells to trigger cytotoxic activities.</li><li>Crp promotes the differentiation of human monocytes toward a pro-inflammatory M1 macrophage phenotype.</li><li>In addition, Crp treatment of M2 macrophages induced</li></ul>
205690	Сгр	21415385	the expression of pro-inflammatory genes and a M1 phenotype. (Demonstrated in human) The major acute phase protein in humans that interacts with CFH of the alternative pathway of complement
103911	CRP	16751408	and C4BP of the classical complement pathway, limiting excessive complement activation CRP induces FCAR surface expression, phagocytosis, and TNF secretion in neutrophils. In addition, CRP physically interacts with FCAR, and induces ERK
103911	CRP	21383176	<ul><li>phosphorylation, cytokine production, and degranulation in mast cells.</li><li>CRP promotes the differentiation of human monocytes toward a pro-inflammatory M1 macrophage phenotype.</li><li>In addition, CRP treatment of M2 macrophages</li></ul>
103911	CRP	21415385	induced the expression of pro-inflammatory genes and a M1 phenotype. Lgmn is an asparagine endopeptidase that removes the majority of the Tlr9 ectodomain, and this catalytic
165744	Lgmn	21402738	cleavage is required for Tlr9 endolysosome signalling in response to DNA. LGMN is an asparagine endopeptidase that removes the majority of the TLR9 ectodomain, and this catalytic
17093	LGMN	21402738	cleavage is required for TLR9 endolysosome signalling in response to DNA. (Demonstrated in murine model) SREBF2 is a key transcriptional regulator of sterol biosynthesis in lipid metabolism, and SREBP2 protein levels in macrophages are negatively regulated by type
9448	SREBF2	21408089	I interferon signalling during viral infection. (Demonstrated in murine model)

133028	Cops5	21403132	Cops5 is required for activation of pro-inflammatory kinases, p38 and ErK, and the down-regulation of Nfe2l2 gene targets. Mice with Cops5 deficiency have lower mortality in polymicrobial sepsis. COPS5 is required for the activation of pro- inflammatory kinases, p38 and ErK, and the down- regulation of NFE2L2 gene targets. Mice with Cops5
24191	COPS5	21403132	deficiency have lower mortality in polymicrobial sepsis. (Demonstrated in murine model) RELB is a NF-kappaB subunit that participates in endotoxin tolerance by repressing pro-inflammatory
57021	RELB	16951372	gene expression. RELB mediates transcription of chemokines like IL8 via activation of AHR and Protein kinase A, and its
57021	RELB	17823304	expression is inhibited by Vitamin D3 analog in DCs. RELB acts as both transcription factor as well as a repressor of NF-kappaB gene expression by forming heterodimers with NFKB1 (p50 subunit) and NFKB2 (p100 subunit) or inhibiting RELA DNA binding
57021	RELB	12657634	<ul> <li>(proo subunit) of minoring KELA DNA onding activity, respectively.</li> <li>RELB sustains NFKBIA (IkappaB alpha) expression during endotoxin tolerance, RelB transcription activation requires binding to the (IkappaB alpha) proximal promoter along with NFKB1 (p50), and is associated with an apparent dimer exchange with</li> </ul>
57021	RELB	19020113	RELA (p65). RELB functions as a dual transcription regulator during LPS tolerance and human severe systemic
57021	RELB	19020113	<ul><li>inflammation (SSI) by activating and repressing innate immunity genes.</li><li>RELB is required for IL17A production in T cell in response to bacterial infection. RELB deficient T cells</li></ul>
57021	RELB	21419662	resulted in a diminished innate immune response to E. coli infection. (Demonstrated in murine model) Xbp1 transcription factor regulates innate immune responses in macrophages via Tlr2 activation and its deficiency results in a much greater bacterial burden in
144745	Xbp1	20351694	mice infected with the Tlr2-activating human intracellular pathogen Francisella tularensis. Xbp1 is an important regulator in ER stress response and may function collaboratively with innate immunity to maintain cellular homeostasis. Xbp1 is essential in poly(I:C)-signalling and ER stress-amplified IFNB
144745	Xbp1	21400498	production in dendritic cells, and the over-expression of Xbp1 synergistically augments the poly(I:C)- induced inflammatory response. XBP1 transcription factor regulates innate immune responses in macrophages via TLR2 activation and its deficiency results in a much greater bacterial burden in mice infected with the TLR2-activating human
3430	XBP1	20351694	intracellular pathogen Francisella tularensis.

			XBP1/ERN1 (IRE1)-mediated signalling plays roles in the coordination of metabolic and immune responses by acting as a regulatory hub, linking endoplasmic reticulum homeostasis with innate immunity and
3430	XBP1	20533428	metabolism. XBP1 interacts with EP300 to augment IFN-beta
3430	XBP1	20660350	induction via a cis-acting enhancer in macrophages under endoplasmic reticulum stress. XBP1 is an important regulator in ER stress response and may function collaboratively with innate immunity to maintain cellular homeostasis. XBP1 is essential in poly(I:C)-signalling and ER stress-amplified IFNB production in dendritic cells, and the over-expression of XBP1 synergistically augments the poly(I:C)-
3430	XBP1	21400498	induced inflammatory response. (Demonstrated in murine model)
75722	NFIL3	19749763	NFIL3 is essential for generation of the natural killer (NK) cell lineage.
			NFIL3 is an IL12B transcriptional inhibitor in macrophages. Interactions of macrophages with the enteric microbiota induce NFIL3 to limit their inflammatory capacity. (Demonstrated in murine
75722	NFIL3	21383239	model) Mbl2 interacts with Fcna and fibrinogen/fibrin to augment the lectin complement pathway, which collaborates with the coagulation system in the first- line host defence against pathogens under conditions
158106	Mbl2	20375621	such as injury and inflammation. Mbl2 treatment inhibits the activity of NFKB and consequently suppresses the production Tnf and Il12 production in human monocytes stimulated with LPS. In addition, Mbl2 was found to bind to Tlr4 and
158106	Mbl2	21383675	attenuate the binding of LPS to cell surfaces. MBL2 is a major recognition molecule of the lectin
74690	MBL2	16105157	pathway of complement. MBL2 binds directly to a wide range of repeating sugar
74690	MBL2	16911830	moieties on microbial surfaces via its lectin domain, resulting in neutralization and opsonization.
74690	MBL2	19840833	MBL2 binds toll-like receptor 4 (TLR4) and modulates cellular responses by altering signals through TLRs. MBL2 treatment inhibits the activity of NFKB and consequently suppresses the production TNF and IL12 production in human monocytes stimulated with LPS.
74690	MBL2	21383675	In addition, MBL2 was found to bind to TLR4 and attenuate the binding of LPS to cell surfaces. Ccr6 mobilizes TCR-alpha/beta+, Ccr6+ innate and adaptive effector T cells in the airway in response to mycobacterial infection. Ccr6 is not required for
135484	Ccr6	21042003	induction of the adaptive antimycobacterial response

135484	Ccr6	21376174	Ccr6 is a chemokine receptor that only binds to a single chemokine ligand, Ccl20. Ccr6 is an important receptor that is involved in regulating mucosal immunity by mediating the recruitment of dendritic cells and APCs to the sites of epithelial inflammation. CCR6 mobilizes TCR-alpha/beta+, CCR6+ innate and adaptive effector T cells in the airway in response to mycobacterial infection. CCR6 is not required for
99015	CCR6	21042003	induction of the adaptive antimycobacterial response
99013	CCR0	21042005	CCR6 is a chemokine receptor that only binds to a single chemokine ligand, CCL20. CCR6 is an important receptor that is involved in regulating mucosal immunity by mediating the recruitment of dendritic cells and APCs to the sites of epithelial
99015	CCR6	21376174	inflammation. Trp63 is a crucial regulator downstream of Tlr3 in Poly(I:C)-induced signalling. Trp63 activates the signalling of both extrinsic and intrinsic apoptosis pathways in endothelial cells through death receptors
150133	Trp63	21367858	and mitochondria. TP63 is a crucial regulator downstream of TLR3 in Poly(I:C)-induced signalling. TP63 activates the signalling of both extrinsic and intrinsic apoptosis pathways in endothelial cells through death receptors
69487	TP63	21367858	and mitochondria. MTOR is the signalling molecule involved in TLR- mediated IFN-alpha production by plasmacytoid
89258	MTOR	18758466	dendritic cells (pDCs). MTOR is an indispensable component of pathogen recognition receptor (PRR) signal pathways that orchestrates the defence program of innate immune
89258	MTOR	18924132	cells. MTOR plays a central role in cell growth and cellular responses to metabolic stress and its activation is essential in TLR2- and TLR4-induced neutrophil activation, as well as in the development and severity
89258	MTOR	19131641	of acute lung injury. MTOR signalling is one major mechanism in a tightly regulated network of intracellular signal pathways including the JAK/STAT system to regulate invasion in human trophoblast cells by secretion of enzymes that remodel the extra-cellular matrix (ECM) such as
89258	MTOR	19331815	MMP2, MMP9, PLAU and SERPINE1. Inhibition of mTOR blocks the anti-inflammatory potency of glucocorticoids both in human monocytes
89258	MTOR	21368289	and myeloid dendritic cells. Cltc functions as a built-in molecular brake that ensures a tight control of basal NFKB activation and gene expression in un-stimulated cells. Defects in Cltc expression could potentially lead to chronic
207019	Cltc	21364927	inflammation disorder.

			CLTC functions as a built-in molecular brake that ensures a tight control of basal NFKB activation and gene expression in un-stimulated cells. Defects in
			CLTC expression could potentially lead to chronic
61767	CLTC	21364927	inflammation disorder. Fgf7 enhances alveolar host defence through GM-CSF-
			stimulated macrophage activation. Intrapulmonary Fgf7 injection enhanced the clearance of E. coli or P.
203973	Fgf7	21343299	aeruginosa via the augmented recruitment, phagocytic activity and oxidant responses of macrophages.
	5		FGF7 enhances alveolar host defence through GM- CSF-stimulated macrophage activation.
			Intrapulmonary FGF7 injection enhanced the clearance of E. coli or P. aeruginosa via the augmented
11473	FGF7	21343299	recruitment, phagocytic activity and oxidant responses
114/3	FOF/	21343299	of macrophages. (Demonstrated in murine model) NFKB1 (p50) is a distinct form of NF-kappaB that
31974	NFKB1	14593105	interacts with STAT3 and cooperates with STAT3 bound to GAS sites.
51771		11095105	NFKB1 is a subunit of the NF-kappaB transcriptional
			regulator complex. NF-kappaB is an inducible transcription factor that regulates the expression of
31974	NFKB1	8152812	numerous genes involved in immune and inflammation
519/4	ΝΓΚΟΙ	8132812	responses and in cellular growth control. NFKB1 enforces specificity of cellular response to
			pathogens by binding to a subset of IRE sequences in IFN-inducible genes. NFKB1 deficiency results in the
			inappropriate production of IFNB in response to
31974	NFKB1	21343618	bacterial DNA sensed by TLR9. (Demonstrated in murine model)
			Cebpe is a member of the CCAAT enhancer binding protein family and is a transcriptional factor regulating
			genes in innate immunity and inflammation. The
162142	Cebpe	21326902	activities of CEBP are regulated via methylation of arginine and lysine side chains.
1021.2	C C C P C		Cebpd is a member of the CCAAT enhancer binding
			protein family and is a transcriptional factor regulating genes in innate immunity and inflammation. The
216094	oleculeID 2160	21326902	activities of CEBP are regulated via methylation of arginine and lysine side chains.
210071		21320702	CEBPE is a member of the CCAAT enhancer binding
			protein family and is a transcriptional factor regulating genes in innate immunity and inflammation. The
3098	CEBPE	21326902	activities of CEBP are regulated via methylation of arginine and lysine side chains.
5070		21520702	CEBPD is involved in TLR8 mediated innate immune response by binding to C/EBP cis-acting elements
281563	CEBPD	20829351	within the TLR8 promoter increasing its transcriptional activity.
			2

CEBPD is a member of the CCAAT enhancer binding protein family and is a transcriptional factor regulating genes in innate immunity and inflammation. The activities of CEBP are regulated via methylation of 21326902 arginine and lysine side chains. Coro2a mediates transcriptional activation of TLR-

responsive genes through a Coro2a-actin-dependent mechanism to remove nuclear receptor co-repressor (NCoR) complexes from the promoters of target genes.

CORO2A mediates transcriptional activation of TLRresponsive genes through a CORO2A-actin-dependent mechanism to remove nuclear receptor co-repressor (NCoR) complexes from the promoters of target genes.

 21331046 (NCoR) complexes from the promoters of target genes. Hmox1, a downstream signalling molecule in the Tlr4 pathway, is necessary for LPS-induced autophagy signalling in macrophages.

HMOX1, a downstream signalling molecule in the TLR4 pathway, is necessary for LPS-induced autophagy signalling in macrophages.

Irak2 regulates cytokine and chemokine mRNA stability and translation in response to LPS stimulation in macrophages. The kinase activity of Irak2 is required for the optimal activation of mitogen activated protein kinase signalling, which regulates cytokine/chemokine production at post-transcriptional levels.

TBKBP117568778A novel component of innate antiviral immunity, it<br/>shares a TBK1-binding domain with AZI2 and TANK<br/>SIAH2 decreases TNF-alpha dependent induction of

281563

145441

78227

168890

5374

178161

56118

61301

62489

64443

64443

65720

66030

CEBPD

Coro2a

CORO2A

Hmox1

HMOX1

Irak2

SIAH2

CD37

SLC15A4

SLC15A4

SYP

21331046

21307647

21291324

17182550

21045126

MAPK8 (JNK) activity and transcriptional activation of NF kappa B by mediating ubiquitination of TRAF2 under stress conditions.

CD37 is important for CLEC7A (Dectin-1) stabilization in APC membranes and controls Dectin-1-mediated IL6 production.

SLC15A4 is an oligopeptide transporter expressed in early endosomes which is involved in transportation of NOD1 ligands.

> SLC15A4, a peptide/histidine transporter in organelle trafficking, is required for the production of proinflammatory cytokines in plasmacytoid dendritic cells upon recognition of viral nucleic acids by endosomal TLR7 or TLR9.

SYP is a protein tyrosine phosphatase that associates with IL6ST and JAK2 in IL-11 signal transduction pathway.

MERTK is a tyrosine protein kinase, it acts with TYRO3 and AXL as pleiotropic inhibitor of the innate immune response in DCs.

66516 MAP2K6 9841871 MAP2K6 activates MAPK14 (p38) MAP kinases.

67727NLRP1217418609NLRP12 is a negative regulator of the NF-ΰB response6772717418609in monocytes.

67727	NLRP12	18280719	NLRP12 negatively regulates non-canonical NF-ΰB pathway by inducing NIK degradation. NLRP12 is an antagonist of toll-like receptor-, tumour
67727	NLRP12	16203735	necrosis factor alpha-, and Mycobacterium tuberculosis-induced pro-inflammatory signals. MMP7 is responsible for cleavage of several mouse
69118	MMP7	19181662	<ul> <li>(Defb1 and Defb2) and human (DEFA1, DEFB1, DEFB4) defensins from pro- to active forms.</li> <li>MMP12 has antimicrobial activity where upon bacterial infection, MMP12 is mobilized to macrophage phagolysosomes and adheres to bacterial cell walls where it disrupts cellular membranes</li> </ul>
69319	MMP12	19536155	resulting in bacterial death.
69883	NLRP2	18056399	NLRP2 is an inhibitor of the NF-kappaB pathway. NLRP2 associates with PYCARD, RIPK2, and CASP1, forming an inflammasome with high proIL-
69883	NLRP2	15030775	1beta-processing activity. NLRP9 is a part of the NLRP (Nucleotide-binding oligomerization domain, Leucine rich Repeat and Pyrin domain containing) family, has a role in apoptosis and
71104	NLRP9	18648497	inflammation. NLRP11 is a part of the NLRP (Nucleotide-binding oligomerization domain, Leucine rich Repeat and Pyrin domain containing) family, has a role in apoptosis and
71154	NLRP11	18648497	inflammation. NLRP13 is a part of the NLRP (Nucleotide-binding oligomerization domain, Leucine rich Repeat and Pyrin domain containing) family, has a role in apoptosis and
71230	NLRP13	18648497	inflammation. NLRP8 is a part of the NLRP (Nucleotide-binding oligomerization domain, Leucine rich Repeat and Pyrin domain containing) family, has a role in apoptosis and
71259	NLRP8	18648497	inflammation. NLRP5 is a part of the NLRP (Nucleotide-binding oligomerization domain, Leucine rich Repeat and Pyrin domain containing) family, has a role in apoptosis and
71285	NLRP5	18648497	inflammation. PSMA7 physically associates with and is involved in the stability of MAVS, which thus provides negative
83912	PSMA7	19734229	regulation of the innate antiviral response against infection by RNA viruses. TNFRSF18 is expressed in several cells and tissues, including T and Natural Killer (NK) cells and is activated by its ligand, GITRL, mainly expressed on Antigen Presenting Cells (APCs) and endothelial cells
84584	TNFRSF18	19760073	<ul><li>and is a modulator of immune response and inflammation.</li><li>CDK9 heterodimerizes with CCNT1 to form the positive transcriptional elongation factor b (P-TEFb) and plays a role in the activation of a subset of NF-</li></ul>
86437	CDK9	18728388	kappaB dependent targets.

			PTAFR is involved in G-protein-independent
95161	PTAFR	11309383	activation of TYK2. C8A is the alpha subunit of complement factor 8 (C8)
			and is one of five components that interact to form the
99144	C8A	12220191	cytolytic membrane attack complex (MAC). SELE is upregulated by TNF alpha during
			inflammatory responses and activated by RELA,
104701	SELE	9006914	NFKB2, JUN and ATF2.
1050 10		• • • • • • • • •	C4A is one of two isotypes of the fourth complement
125942	C4A	2650988	component.
125042	C14	1547(020	C4A is a soluble complement anaphylatoxin that
125942	C4A	15476920	greatly controls the local pro-inflammatory response.
			Mbl1 interacts with Fcna and fibrinogen/fibrin to
			augment the lectin complement pathway, which collaborates with the coagulation system in the first-
			line host defence against pathogens under conditions
151802	Mbl1	20375621	such as injury and inflammation.
131802	WIDII	20373021	DEFA3 transcribes a large amount cationic alpha-
			defensin peptides, HNP-3, in human neutrophils, these
			alpha-defensing have multiple functions in the immune
296590	DEFA3	19024344	system.
			Apcs is a key negative regulator for innate immune
			responses to DNA and may be partly responsible for
			the insufficient immune responses after DNA
			vaccination in humans. Murine Apcs exhibited a
			similar, albeit very weak, activity.
205704	Apcs	21278351	
			Cfh, a complement regulatory factor, interacts with
			host cell surfaces as well as C3d part of C3b and plays
			a major in the distinguishing host from non-host
			surfaces during the alternatively activated complement
196803	Cfh	21285368	pathway.
170005	CIII	21203300	PTPN2 pays a novel role in the regulation of type 1
			interferon-stimulated gene expression in cells
1165	PTPN2	14600148	previously desensitized to Type 1 interferons.
			PTPN2 is a nuclear protein tyrosine phosphatase, a
			potential negative regulator of the PRL-mediated
1165	PTPN2	11773439	signalling pathway.
			IFNGR2 associates with JAK2 and this is required for
2140	IFNGR2	9001223	IFN-gamma signalling.
			IFNRG2 complex is activated by STAT1, leading to
			the transcription of a significant portion of IFN-gamma
			induced genes, many of which are responsible for the
2140	IFNGR2	16467883	induction of an apoptotic state in response to IFN-
2140	IF NUK2	1040/883	gamma. RCAN1 (DSCR1) short isoform positively modulates
			IL-1R-mediated signalling pathways by regulating
2768	RCAN1	19716405	TOLLIP/IRAK1/TRAF6 complex formation.
2700	NUANI	17/10-03	

			Inhibitory SMAD7, a direct target gene for transforming growth factor-beta (TGF-beta), mediates TGF-beta1-induced apoptosis in several cell types and acts as a scaffolding protein to facilitate TAK1- and
3195	SMAD7	12589052	MKK3-mediated activation of p38. SMAD7 acts as a critical mediator for effective TGF- beta1-mediated suppression of IL-1R/TLR signalling, by simultaneous binding to discrete regions of Pellino-
3195	SMAD7	20171181	1. MALT1 forms a complex with CARM1 and BCL10 to
3930	MALT1	18192506	MALTI ionis a complex with CARWIT and BCLTO to activate NF-kappaB. MALT1 is a paracaspase that has arginine-directed proteolytic activity. MALT1 cleaves TNFAIP3, a dual ubiquitin-editing enzyme involved in termination of NF-kappaB signalling, inducing cytosolic release of
3930	MALT1	20804738	TNFAIP3 and dampening its inhibitory function. ABCG1 regulates innate immunity in a tissue-selective manner and Abcg1(-/-) mice have an enhanced pulmonary host defence response driven predominantly
4231	ABCG1	20395559	by hematopoietic cells. SOCS6 associates with KIT and regulates KIT receptor
4847	SOCS6	14707129	signalling and leads to MAPK activation. CSF2RB (IL3RB) is tyrosine phosphorylated by the simultaneous activation of both JAK1 and JAK2 fusion proteins, but not either one alone. Phosphorylated CSF2RB then induces the activation of downstream signaling molecules, including STAT5, AKT, and MAPK, and the conferring of factor-independent
6178	CSF2RB	15988755	growth to IL-3-dependent Ba/F3 cells IL2RB is phosphorylated by JAK1and different IL2RB tyrosines couple to at least two signalling pathways
6418	IL2RB	8700888	<ul><li>(JAK-STAT and SHC-coupled) and synergistically mediate IL2-induced proliferation.</li><li>IL2RB physically associates with JAK1 suggesting that regulation of JAK1 may be linked to IL2 induced</li></ul>
6418	IL2RB	8041779	signal transduction. LGASL2 binds to T cells in a beta-galactoside-specific manner and acts as pro-apoptotic effector for activated
6696	LGALS2	15356130	T cells. APOBEC3A has deaminase activity in monocytes and
7865	APOBEC3A	20615867	macrophages and this induces high levels of TC- specific deaminase activity in IFN-alpha signalling. MAP3K7IP1 induces MAP3K7
8281	MAP3K7IP1	15590691	autophosphorylation/activation and inhibits MAP3K7 interaction with the IKK signalosome. MAP3K7IP1 functions as an activator of the MAP3K7 (TAK1) in TGF-beta signal transduction.
8281	MAP3K7IP1	8638164	

			MAP3K&IP1 participates in a SAPK2a/p38alpha- mediated feedback control of MAP3K7, which not only limits the activation of SAPK2a/p38alpha but synchronizes its activity with other signalling pathways
8281	MAP3K7IP1	14592977	that lie downstream of MAP3K7 (JNK and IKK). MAP3K7IP1 modulates intracellular localization of
8281	MAP3K7IP1	16407200	MAPK14 (p38) and downstream signalling. MLST8 regulates TNF-alpha-induced NF-kappaB signalling by directly inhibiting the activation of
10142	MLST8	18755269	IkappaB kinase. TCEB2 is a part of Elongin BC complex, is a component of a multiprotein SOCS1 complex that attenuates JAK/STAT signaling by binding to JAK2 and inhibiting JAK2 kinase and by interacting with SOCS box, the Elongin BC complex can increase expression of the SOCS1 protein by inhibiting its
10956	TCEB2	9918119	degradation. TCEB2, as a part of the Elongin B and C complex, is
10956	TCEB2	10051596	bound by SOCS protein for proteasomal degradation. MEFV inhibits the formation of CASP8, PYCARD and NLRC4 inflammasome formation by competing
11729	MEFV	126461680	with CASP8 to bind to PYCARD.
11729	MEFV	12615073	MEFV inhibits binding of NLRP3 with PYCARD. PYCARD-MEFV pyroptosome is induced by PSTPIP1
11729	MEFV	17964261	in a pyrin-dependent manner. MEFV homodimerization is important for its ability to induce PYCARD oligomerization and CASP1
11729	MEFV	17964261	activation. MEFV gene is bound constitutively by CEBPB, while RELA binds to the MEFV gene upon TNFA
11729	MEFV	14514692	stimulation. PTK2B mediates the JAK-dependent activation of
13899	PTK2B	10228162	MAPK and STAT1 in interferon (IFN)-gamma, but not IFN-alpha signalling. PTK2B amplifies epidermal growth factor receptor (EGFR) and SRC-induced STAT3 activation, implicating PTK2B activation as a potential co-
13899	PTK2B	14963038	mediator in triggering STAT3-induced oncogenesis. C1q/TNF-related protein-3 (AMACR;C1QTNF3), a protein secreted by adipocytes (adipokine), inhibits three basic and common pro-inflammatory pathways involved in obesity and type 2 diabetes mellitus (adipo- inflammation) by acting as an endogenous LPS
15015	MACR;C1QTN	20739398	antagonist of the adipose tissue. SCARF1 mediates host defence against Cryptococcus neoformans and Candida albicans through cytokine production and is required for macrophage binding to
15121	SCARF1	19237602	C. neoformans to control the infection in mice.

PTPN6 (SHP-1) normally functions to antagonize the IL-2 signal transduction pathway and human T-lymphotropic virus type I (HTLV-I) infection and oncogenic transformation can lead to loss of SHP-1 expression, resulting in constitutive activation of IL-2 regulated T cell responses

15719	PTPN6	9520455	regulated T cell responses.
			PTPN6 reversibly associates with the IFN-alpha
			receptor complex upon IFN stimulation and selectively
			regulates distinct components of JAK/STAT signal
15719	PTPN6	8524272	transduction pathways.
			PTPN6 specifically downregulates MAP3K7 (TAK1)
15710		1202020	through dephosphorylation, suppressing inflammatory
15719	PTPN6	17079228	responses via TAK1 signalling pathways.
15710	DTDNC	19201054	PTPN6 promotes TLR- and RIG-I-activated production
15719	PTPN6	18391954	of type I interferon by inhibiting the kinase IRAK1.
			PTPN6 tyrosine phosphatase activity plays a critical
15719	PTPN6	20145200	role in induction of IL2B production in macrophages in response to TLR ligands.
13/19	I II NU	20143200	CLEC4C (BDCA-2) signalling inhibits TLR-9-
			agonist-induced plasmacytoid dendritic cell activation
			(through the inhibition of CD86 and CD40) and
16472	CLEC4C	20673884	antigen presentation.
101/2	CLECIC	20072001	SIGIRR is a negative modulator of TLR-IL-1R
			signalling where it binds to the TLR-IL-1R signalling
16551	SIGIRR	12925853	components in a ligand-dependent way.
			SIGIRR controls Th17 cell expansion and effector
			function through the IL-1-induced mTOR signalling
16551	SIGIRR	20060329	pathway.
			SIGIRR regulates innate responses in differentiated
			human intestinal epithelial cells (IECs), modulating
			epithelial involvement in infectious and inflammatory
16551	SIGIRR	20130217	bowel diseases.
			SIGIRR can inhibit Toll-like receptor (TLR) 4, 5, and
			9-mediated immune responses by attenuating
			production of the inflammatory mediators IL-6 and
			TNF-alpha and this attenuation was not the result of
			decreased expression of TLR4, 5 or 9, but rather a
16551	SIGIRR	20364327	sequestration of MYD88 to the TLRs.
			HRAS participates in CpG oligodeoxynucleotide
			signalling through association with TLR9 and
16070		12067410	promotion of IRAK1/TRAF6 complex formation in
16878	HRAS	12867418	macrophages.
			HRAS participates in the activation of MAPK1 by IL1A (IL-1) through association with IRAK1, IRAK2,
16878	HRAS	11744690	TRAF6 and MAP3K7.
100/0	IIIAS	11/44090	HRAS promotes virus spread by suppressing viral
			RNA-induced IFN-beta production through negative
			KINA-Induced IFIN-beta production through negative

16878HRAS20501842RNA-induced IFN-beta production through negative<br/>regulation of RIG-I signalling.<br/>C19orf29 (Cactin) is a novel negative regulator of TLR<br/>signalling that targets the MHC class III protein<br/>I{kappa}B like (I{kappa}BL) and inhibits NFkappaB17610C19orf2920829348and IRF signalling pathways.

17892	С9	16189651	C9, Complement factor nine, is one of five components that interact to form the cytolytic membrane attack complex (MAC), composed of a C5B-C8 complex attached to a transmembrane C9 oligomer. PIAS4 is a member of the Protein Inhibitor of Activated STAT protein family, which regulate innate immune response by controlling transcription induced by TLR, RLR and JAK/STAT signalling pathways. PIAS4 specifically negatively regulate both IFN transcription and IFN stimulated gene expression through multiple mechanisms utilizing the function of different domains.
18183	PIAS4	21199872	Pias4 is a member of the Protein Inhibitor of Activated STAT protein family, which regulate innate immune response by controlling transcription induced by Tlr, Rlr and Jak/Stat signalling pathways. Pias4 specifically negatively regulate both Ifn transcription and Ifn stimulated gene expression through multiple mechanisms utilizing the function of different domains.
177127	Pias4	21199872	MAP2K2 (MEK2)/PI3CD is a novel IFN-beta triggered signalling cascade that regulates secreted IL- 1Ra (sIL-1Ra) expression in monocytes and this provides a rationale for an alternative, interferon (IFN)- beta-mediated pathway to induce/enhance sIL-1Ra
18293	MAP2K2	20837746	production, dampening inflammation. SMAD6 is a critical mediator of the TGF-beta-BMP
18304	SMAD6	16951688	pathway that mediates anti-inflammatory activity and negatively regulates IL-1R-Toll-like receptor signals. SMAD6 acts a critical mediator for effective TGF- beta1-mediated suppression of IL-1R/TLR signalling, by simultaneous binding to discrete regions of Pellino-
18304	SMAD6	20171181	1. STAP2 acts as an endogenous negative regulator of EBV LMP1-mediated signalling through TRAF3 and
18753	STAP2	18573890	TRADD. TOLLIP is a negative regulator of toll-like receptor (TLR) mediated signalling which serves to limit the production of pro-inflammatory mediators during
19140	TOLLIP	11751856	inflammation and infection. TOLLIP over-expression inhibits activation of NF-
19140	TOLLIP	11751856	kappaB in response to IL1, the TLR2 and TLR4 ligands. TOLLIP/IRAK1/TRAF6 complex formation is positively regulated by RCAN1 (DSCR1) short
19140	TOLLIP	19716405	isoform, resulting in the modulation of IL-1R-mediated signalling pathways. ARRB2 acts to limit JNK/ERK activation and survival
19263	ARRB2	19783052	in macrophages and is required for basal and TLR- inducible complement C1q expression.

ARRB2 regulates TLR4-mediated apoptotic signalling through GSK3B where ARRB2 represents an inhibitory effect on the TLR4-mediated apoptotic cascade, through controlling the homeostasis of 19263 ARRB2 20497256 activation and inactivation of GSK3B. RE-1 silencing transcription factor (REST) is part of a repressor complex, along with key components that include histone deacetylase (HDAC) 1 or 2, corepressor of REST (CoREST), and lysine-specific demethylase (LSD) The 1. HDAC/CoREST/REST/LSD1 repressor complex is a 20380 REST 20798038 significant component of host innate immunity. HSP90AA1 (HSP90) positively regulates NOD1 20861 HSP90AA1 17420470 activation. HSP90AA1 regulates the stability of transforming growth factor beta-activated kinase 1 (TAK1) in interleukin-1beta (IL1B)-induced cell signalling and IL1B-induced signalling by interacting with and maintaining the stability of TAK1, suggesting that HSP90AA1 might act as the chaperone of TAK1 in immune and inflammatory responses related with IL1A 20861 HSP90AA1 18950863 (IL-1) signal cascades. Co-repressor of REST (RCOR1) is part of a repressor complex, along with key components that include histone deacetylase 1 or 2, RE-1 silencing transcription factor (REST), and lysine-specific demethylase (LSD) 1. The HDAC/RCOR1/REST/LSD1 repressor complex 21216 RCOR1 20798038 is a significant component of host innate immunity. KAT2B is a histone acetylase and a phorbol esterinducible co-activator of the interferon (IFN) regulatory factor (IRF) proteins which contributes to the 21805 KAT2B 10022868 establishment of type I IFN responsiveness. NLRP1 forms а biochemical complex, or 21836 NLRP1 18280719 inflammasome, with NALP1, CARD7, and DEFCAP. NLRP1 forms inflammasome upon cellular infection by Toxoplasma gondii - this process is critical in mediating the innate immune response to T. gondii infection and pathogenesis. 21836 NLRP1 21098108 THRB signalling plays a role in modulating dendritic cell (DC) physiology and has immunoregulatory effects. THRB contains an NF-kappaB consensus site in its promoter region that controls its expression, which in turn signals DCs to promote maturation and function via an Akt-dependent, but PI3K independent 22332 20018842 THRB pathway. MTA1 binds to the MYD88 promoter to regulate LPSinduced NFkappa B signaling via MYD88-dependent signalling in murine macrophages.

23411

MTA1

20699220

			Mta1 binds to the Myd88 promoter to regulate LPS- induced NFkappa B signaling via Myd88-dependent
175870	Mtal	20699220	signalling in murine macrophages.
23908	CCR4	18624303	<ul><li>CCR4 modulates TLR9-mediated innate immunity and signalling.</li><li>IL8 is produced by airway epithelial cells in response to invading bacteria and mediates airway epithelial</li></ul>
23954	IL8	17220369	defence against bacterial infection via the DUOX1- TACE-TGF-alpha-EGFR signalling pathway. CD36 mediates host defence against Cryptococcus neoformans and Candida albicans through cytokine
24023	CD36	19237602	production. CD36 ectodomain binds negatively charged diacylglycerol ligands and CD36, along with CD14,
24023	CD36	19847289	has a non-redundant role for loading ligands onto TLR2 in the plasma-membrane. CD36 is a selective and non-redundant sensor of microhial disculateoridae that signal via the TLP2/6
24023	CD36	15690042	microbial diacylglycerides that signal via the TLR2/6 heterodimer. CD36-TLR4-TLR6 activation is a common molecular
24023	CD36	20037584	mechanism by which atherogenic lipids and amyloid- beta stimulate sterile inflammation. CD180 is an accessory molecule for TLR2, forming
24968	CD180	19154986	part of the receptor complex for innate immune recognition of mycobacterial lipoproteins. CD180 and its helper molecule, LY86, interact directly with the TLP4 ciercelling complex inhibiting its
24968	CD180	15852007	with the TLR4 signalling complex, inhibiting its ability to bind microbial ligands. CD180 dramatically enhances CpG DNA-induced proliferation/survival by naive B cells but not by memory B cell. This enhancement that is mediated by CD180-induced TLR9 upregulation, leading to Akt
24968	CD180	20133206	activation and sustained NF-kappaB activation. BCL2A1 negatively regulates autophagy and expression of BCL2A1 in Mycobacterium tuberculosis infected macrophages provides the bacteria a survival strategy to overcome host defences.
25591	BCL2A1	21167304	TCEB1 is a part of Elongin BC complex, and is a component of a multiprotein SOCS1 complex that attenuates JAK/STAT signalling by binding to JAK2 and inhibiting JAK2 kinase and by interacting with SOCS box, the Elongin BC complex can increase
25766	TCEB1	9918119	expression of the SOCS1 protein by inhibiting its degradation. IRAK4 is required for interleukin-1 receptor/toll-like
28022	IRAK4	18794297	receptor-induced MAP3K7 (TAK1)-dependent NF- kappaB activation. IRAK4 and IRAK1 play key roles in a signalling
28022	IRAK4	17997719	pathway by which bacterial infection or IL-1 trigger the production of inflammatory mediators.

			IRAK4 is indispensable for the responses of animals and cultured cells to IL-1 and ligands that stimulate
28022	IRAK4	11923871	various Toll-like receptors (TLRs). IRAK4 plays a critical role in IL-1R and TLR
28022	IRAK4	12682231	signalling cascades and is an essential component of the IL-18 signalling cascade. IRAK4 activation is impaired during endotoxin
			tolerization; a process which impairs the production of LPS-induced pro-inflammatory cytokines without inhibition expression of anti-inflammatory or anti-microbial mediators.
28022	IRAK4	21220427	
			ITGAM (CD11b integrin) is activated via Toll-like receptors (TLRs) and engages in crosstalk with the MYD88 and TICAM1 (TRIF) pathways inhibiting
28139	ITGAM	20639876	TLR signalling in innate immune responses. ITGAM :: ITGB2 is the principal leukocyte receptor
			involved in the recognition of the fungus Candida albicans. Recognition of Pra1p protein of C. albicans by ITGAM :: ITGB2 plays a pivotal role in determining fungal virulence, and host response/protection against C. albicans infection.
28139	ITGAM	21245270	(Demonstrated in murine model)
20137	HOAM	21243270	VDR, vitamin D receptor, deletion leads to reduced
			level of NFKBIA protein through protein translation, protein-protein interaction, and post-translational
29237	VDR	19931640	modification.
30112	SIAH1	11742346	SIAH1 is a ubiquitin ligase structurally related to TRAF and modulates TNF-alpha signalling.
			LPCAT2 is highly expressed in inflammatory cells and is activated by lipopolysaccharide (LPS) treatment through Toll-like receptor 4 (TLR4) and LPCAT2 phosphorylation through LPS-TLR4 signalling may directly depend on MAPK-activated protein kinase 2
31324	LPCAT2	20663880	(MAPKAPK2). ACHE (acetylcholinesterase) expression is induced by
32513	ACHE	18385943	hydrogen peroxide (H2O2) via the JNK/AP1/ ATF2 signalling pathway.
			PALM3 is a LPS inducible gene that functions as an adaptor protein for TLR4 signalling. PALM3 interacts
22017	(alamiaD 2294	21197075	with SIGIRR to negatively regulate TLR signalling.
32847	IoleculeID 3284	21187075	TRAIP is a SYK antagonist in TNF signaling -
35609	TRAIP	19151749	overexpression of TRAIP sensitize cells to TNF- induced apoptosis.
			TRADD is recruited to MAVS and orchestrated complex formation with TRAF3 and TANK and with FADD and RIPK1, leading to the activation of IRF3
36010	TRADD	18439848	and NF-kappaB-TRADD is not only essential in TNFR1 signalling but also in RIG-I antiviral pathway.

			TNIP3 is a novel lipopolysaccharide-inducible inhibitor of NF-kappaB activation, binds to A20 and inhibits NF-kappaB activation induced by tumour necrosis factor, interleukin-1, and 12-O-
36076	TNIP3	17088249	tetradecanoylphorbol-13-acetate. MAP3K12 (DLK) and its downstream kinases contribute to the finely tuned regulation of CREB- dependent effects. MAP3K12 inhibits CREB activity
36895	MAP3K12	20940047	by affecting the interaction of CREB with its second co-activator TORC. PTK2 induces KLF8 expression in human ovarian
37147	PTK2	18353772	cancer cells by activating the PI3K-Akt signalling pathway. CAV1 is an important component of the innate host immune response to the majority of non-cytotoxic strains of P. aeruginosa by promoting bacterial
37314	CAV1	19949109	clearance during acute pneumonia and chronic colonization. CAV1 is a scaffolding protein of caveolae that plays an important role in host defence and inflammation and
37314	CAV1	20304961	CAV1 deficiency dampens Toll-like receptor 4 signalling through NOS3 (eNOS) activation. ELMOD2 regulates the TLR3 signalling pathway where silencing of ELMOD2 in human macrophages
38838	ELMOD2	19966137	inhibited TLR3-dependent expression of type I and type III interferon genes. RNF41 is an E3 ligase that can negatively regulate MyD88-dependent production of pro-inflammatory
40313	RNF41	19483718	cytokines but can promote TRIF-dependent production of type I interferon. SIRPA negatively regulates TLR4 or TLR3 dependent
40694	SIRPA	18233962	cytokine production through inhibition of NF-kappaB dependent signalling. SIRPA down-regulation is lipopolysaccharide (LPS)
40694	SIRPA	17954568	inducible and contributes to innate immune activation in macrophages. BCAR1 acts as a primary force sensor, transducing
42468	BCAR1	17129785	force into mechanical extension and thereby priming phosphorylation and activation of downstream signalling whereby tyrosine phosphorylation of BCRA1 in a cytoskeletal complex is involved in force- dependent activation of the small GTPase RAP1A.
			Nuclear ZMYND11 is recruited by oligomerized, cytoplasmic TICAM1 to enhance NF-kappaB activation and type-I-interferon induction. ZMYND11 harbours dual modes of cytoplasmic NF-kappaB
43779	ZMYND11	19795416	regulation, positively in the TICAM1 pathway and negatively in the PDLIM7 (LMP1) pathway. ZMYND11 cooperates with TRAF3 in the regulation
43779	ZMYND11	20138174	of Epstein-Barr virus-derived LMP1/CTAR1-induced NF-kappaB activation.

CD22 (Siglec-2), like other sialic acid-binding immunoglobulin-like lectins (siglecs), is predominately expressed on immune cells. CD22 exhibits hallmarks of clathrin-mediated endocytosis and traffics to recycling compartment, reflecting its role in cell signalling and innate immunity.

CD22 expression and function are differentially regulated in B-1 and conventional B-2 cells, which are implicated in innate and adaptive immunity, respectively.

CD274 expression in macrophages is induced upon exposure to HIV virions and TLR stimulation. In addition, IL10 also up-regulates CD274 expression.

PDCD1LG2 expression in macrophages is induced upon exposure to HIV virions, and is transcriptionally down-regulated by IL10.

CCR7 is a chemokine receptor expressed on the surfaces of T cells, B cells, and mature dendritic cells that controls cell migration in response to the cognate ligands CCL19 and CCL21. CCR7 deficiency results in a heightened pro-inflammatory environment in response to acute pulmonary P. aeruginosa infection and contributes to more efficient clearance.

47847 SMARCE1 17669635 SMARCE1 is a transcriptional modulator that is known to repress viral replication.

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CD22

**CD22** 

CD274

PDCD1LG2

CCR7

NKIRAS2

DHX58

DHX58

DHX58

BECN1

NUMBL

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16116171

SMARCE1 is associated with up-regulation of<br/>proapoptotic genes, including the tumour suppressorSMARCE116135788familial cylindromatosis (CYLD).

NKIRAS2 is a key inhibitor of NF-kappaB signalling and estradiol represses NF-kappaB activation through the induction of NKIRAS2.

DHX58 (LGP2) helicase is related to DDX58 (RIG-I) and IFIH1 (MDA5) but lacks caspase activation and recruitment domains (CARDs), and functions as a negative regulator of innate host defence.

DHX58 lacks the caspase recruitment domain (CARD) homology and functions as a negative regulator by interfering with the recognition of viral RNA by DDX58 (RIG-I) and IFIH1 (MDA5).

DHX58 functions as a negative regulator of antiviral innate immunity by interfering with the recognition of viral RNA by DDX58 (RIG-I) and IFIH1 (MDA5).

SOCS2SOCS2 can enhance IL-2 and IL-3 signalling byaccelerating SOCS3 degradation.

BECN1 is recruited into the mouse Myd88 and Ticam TLR-signalling complex thereby reducing its binding 18772134 to Bcl2, leading to autophagy.

NUMBL interacts with TAB2 (MAP3K7IP2) and inhibits TNF alpha and IL-1Beta-induced NF-kappaB 18299187 activation.

			CCDC88A, an activator of GNAI3, plays a key role in regulating autophagy; the dynamic interplay between GNAI3, GPSM1 and CCDC88A determines whether autophagy is promoted or inhibited. When stimulated by growth factors, CCDC88A disrupts the GNAI3::GPSM1 complex, subsequently enhancing anti-autophagy signalling pathways and inhibits autophagy by activating GNAI3.
52430	CCDC88A	21209316	RPS19 interacts with macrophage migration inhibitory
53332	RPS19	19155217	factor (MIF) and attenuates its pro-inflammatory function by inhibiting the MIF-CD74 interaction and MIF triggered adhesion of monocytes. CAMK2A promotes TLR-triggered pro-inflammatory cytokine and type-I-IFN production by directly binding and activating MAP3K7 and IRF3 in macrophages. (Demonstrated in murine model)
53452	CAMK2A	18818394	Camk2a promotes Tlr-triggered pro-inflammatory
150657	Camk2a	18818394	cytokine and type-I-IFN production by directly binding and activating Map3k7 and Irf3 in mouse macrophages. IFNA2 controls chemotaxis by regulating the CXC
54199	IFNA2	18729739	receptor ligand interaction between CXCL10 and CXCR3A.
54199	IFNA2	18027911	IFNA2 interacts differentially with IFNAR2, and influences IFNA1 interaction with IFNAR2. IFNA2 induces the transcription of HIF1A in human endothelial cells though a JAK-ISGF3 pathway under
54199	IFNA2	18606657	normoxic conditions, and this response contributes to the anti-proliferative activity of this cytokine. IFNA1 activates NF-kappaB in JAK1-deficient cells through a TYK2-dependent pathway where for the IFN signalling pathway leading to STAT activation, both JAK1 and TYK2 are essential, but NF-kappaB
54276	IFNA1	15883164	activation requires only TYK2. IFNA1 induces the human anti-inflammatory cytokine
54276	IFNA1	12817009	IL-10 gene via a module consisting of interdependent IRF1 and STAT3 motifs. IFNA1 sensitizes cells to microbial recognition by up- regulating the expression of several TLRs as well as
54276	IFNA1	15699120	adapter molecules and kinases involved in TLR signalling. IFNA1 is a type I interferon (IFN) that binds to the IFN
54276	IFNA1	9737881	receptor (IFNAR), composed of two transmembrane polypeptides, IFNAR1 and IFNAR2;IL10RB. MAP3K14 associates with DDX58 (RIG-I) and its downstream adaptor, mitochondrial antiviral signaling (MAVS) and the MAP3K14-DDX58 signalling pathway induces RELA release from NFKB2
54829	MAP3K14	18550535	complexes in response to respiratory syncytial virus (RSV) infection.

MAP3K14 binds to NOD2 and mediates induction of specific changes induced by NOD2 activator muramyl dipeptide (MDP) and this occurs in settings where both the NOD2 and TLR4 pathways are activated by their respective agonists.

ANXA4 differentially modulates the NF-kappaB signalling pathway and this is dependent on its interactions with NFKB1 and the intracellular Ca(2+) ion level.

CYBB (NOX2) functions in reactive oxygen species (ROS) generation to induce TNF-alpha-mediated host cell apoptosis and it function in sensing of persistent intracellular pathogens for subsequent induction of host cell apoptosis as a second line of defence.

CYBB is an NADPH oxidases which plays a central role in microbial killing by phagocytes through the generation of reactive oxygen species (ROS) and CYBB-generated ROS are necessary for LC3 recruitment to phagosomes, coupling oxidative and non-oxidative killing activities of the CYBB NADPH

acting as a negative regulator of Toll-like receptor signalling.

19339495 **CYBB** oxidase in phagocytes through autophagy. 55775 CYBB and reactive oxygen species (ROS) are required for the host cell to trigger an efficient RIG-I-mediated IRF-3 activation and downstream antiviral IFNbeta and IFIT1 gene expression and CYBB is critical for the expression of the central mitochondria-associated 55775 **CYBB** 20532218 adaptor MAVS. BCL3 is critically involved in lung defence against bacteria Klebsiella Gram-negative pneumoniae, modulating functions of several cells to facilitate efficient clearance of bacteria. Loss of BCL3 incurred dramatic cytokine imbalance in the lungs, failure to clear bacteria and increased susceptibility to K. pneumoniae pneumonia. (Demonstrated in murine model) BCL3 21228348 56568 RIPK1 ubiquitination on Lys377 is required for tumour necrosis factor (TNF-alpha) induced NF-kappaB 57326 RIPK1 16543241 activation. RIPK1 is a death domain kinase that is one of the critical components involved in mediating DNA 57326 RIPK1 16825191 damage-induced, p53-independent cell death. TRAFD1 deficiency reveals its negative regulatory role in the TLR and RIG-I-like helicases signalling pathway and was found to interact with TRIF, IPS-1, TRAF3 57921 TRAFD1 18849341 and TRAF6 in mice TRAFD1 is an interferon and LPS inducible gene 57921 TRAFD1 16221674

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MAP3K14

ANXA4

**CYBB** 

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ITGA3 along with ITGB1 is a novel regulator for the recognition of bacterial lipopeptides. ITGA3/ITGB1 integrin regulates endosomal Toll-like receptor (TLR)-2/TLR1 signalling, serving as a mechanism for modulating inflammatory responses.

IRF2BP1 is a transcriptional co-repressor of IRF2 and, through physically association, enhances JDP2 polyubiquitination. IRF2BP1 has also been shown to repress ATF2-mediated transcriptional activation from 18671972 a CRE-containing promoter.

> PIK3CA and PIK3CB isoforms of class IA phosphatidylinositol 3-kinase (PI3K) are both required for the pro-inflammatory response to flagellin.

LY86 and CD180 interact directly with the TLR4 signalling complex, inhibiting its ability to bind 15852007 microbial ligand.

> RANBP9 is located in the Microtubule-Organizing Center it contains protein-interaction motifs, a cytoskeletal-binding domain, and multiple canonical docking sites for signalling intermediates. RANBP9 acts as an scaffolding protein and is important for maintaining cellular functions in the immune and nervous system.

NUP153 and NUP214 are nucleoporins that control the nucleo-cytoplasmic shuttling and, along with XPO1dependent nuclear export, the subcellular distribution 15210729 of latent STAT1.

> OTUD5 is a deubiquitinase that regulates Type I interferon (IFN) production where OTUD5 selectively cleaves the lysine-63-linked polyubiquitin chains on TRAF3, resulting in its dissociation from the downstream signalling complex containing TBK1.

MAP3K8 negatively regulates interferon-beta 19667062 production in macrophages and myeloid dendritic cells. MAP3K8 is essential for IL-1beta production from both macrophages and dendritic cells and an important mediator for collaboration of pattern recognition receptors with danger-associated molecular patterns (DAMPs) to induce TNF and IL-1beta production and optimal host defence. 19933865

> MAP3K8 IL23A regulates expression in lipopolysaccharide (LPS)-stimulated macrophages through extracellular signal-regulated kinase (ERK) activation.

MAP3K8 is a MEK kinase that is require for the activation of MAP kinases in myeloid cells following TLR and TNF receptor stimulation. MAP3K8 is critical for production of the pro-inflammatory cytokine TNF during inflammatory responses.

66703 MAP3K8 21135874

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ITGA3

IRF2BP1

PIK3CB

LY86

RANBP9

NUP153

OTUD5

MAP3K8

MAP3K8

MAP3K8

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			LILRA4 couples with a signalling adapter to activate a prominent immunoreceptor tyrosine-based activation motif (ITAM)-mediated signalling pathway in human
68562	LILRA4	20193018	plasmacytoid dendritic cells. BIRC3 is required for proper RIPK1 polyubiquitination and NF-kappaB activation upon TNF (TNF-alpha) treatment and regulates TNF-
69045	BIRC3	18697935	mediated NF-kappaB activation by binding to TNFRSF1A (TNFR1). BIRC3 and BIRC2 are required for innate immunity signalling by the pattern recognition receptors NOD1
69045	BIRC3	19464198	and NOD2. PROCR confers anti-inflammatory properties when bound by activated protein C and when PROCR (EPCR) is blocked, F. tularensis loses the ability to suppress the pro-inflammatory response of endothelial
69466	PROCR	20543103	cells. Rat CASP12 binds to Human RIPK2 and displaces Human TRAF6 from a complex in human cells,
69532	CASP12	18329614	<ul><li>inhibiting its ubiquitin ligase activity, and blunting NF-kappaB activation.</li><li>CASP12 deficiency enhances production of antimicrobial peptides, cytokines, and chemokines dependent on bacterial type III secretion and the Nod</li></ul>
69532	CASP12	18329614	pathway. CARD18 is an inhibitor of LPS-induced IL1B (IL-1
69714	CARD18	11051551	beta) generation by preventing RIPK2-mediated oligomerization and auto-activation of CASP1. CARD18 is an intracellular regulator of CASP1 (caspase-1) activation and plays a role in the regulation of IL1B (IL-1beta) secretion and NF-kappaB activation
69714	CARD18	11536016	during the pro-inflammatory cytokine response. SOCS3 inhibits the JAK/STAT pathway and act as a
70676	SOCS3	10882725	negative regulator of fetal liver erythropoiesis (EPO) by binding the JAK2 and EPO receptor, respectively. SOCS3 is rapidly induced by IL-2 in T cells where it acts to inhibit IL-2 responses in a classical negative feedback loop by suppressing STAT5 phosphorylation
70676	SOCS3	10373548	and lymphocyte proliferation. SOCS3 protein is a potent endogenous inhibitor of Janus kinase (JAK)/STAT signalling and influenza A virus inhibits type I IFN signalling via NF-kappaB-
70676	SOCS3	18989459	dependent induction of SOCS3 expression. SOCS3 is a molecular inhibitor of IFN signalling and SOCS3 expression, induced by stimuli present in the Human immunodeficiency virus (HIV)-1-infected brain, such as transactivator of transcription, inhibits antiviral IFN-beta signalling to enhance HIV-1 replication in macrophages, allowing HIV-1 to evade the protective innate immune response within the
70676	SOCS3	20631305	central nervous system.

			CYTIP co-localizes with SNX27 at the early endosomal compartment of lymphocytes suggesting a
72074	CVTD	17577500	role for this interaction in endocytic trafficking and/or
72874	CYTIP	17577583	signalling. MAFB binds to IRF3, impairing the recruitment of co-
75124	MAFB	20581830	activators to IRF3 and antagonizing antiviral responses. Sirt1 is a histone deacetylase enzyme that has been found to accumulate at the promoters of Tnfa and Il1b in response to Tlr4 signalling. Sirt1 promotes termination of NFKB-dependent transcription and recruits Relb to assemble transcription repressor complex that generates endotoxin tolerance.
159441	Sirt1	21245135	SIRT1 is a histone deacetylase enzyme that has been found to accumulate at the promoters of TNFA and IL1B in response to TLR4 signalling. SIRT1 promotes termination of NFKB-dependent transcription and recruits RELB to assemble transcription repressor complex that generates endotoxin tolerance.
75847 75863	SIRT1 NFE2L2	21245135 21220332	NFE2L2 plays an important role in TLR4-mediated autophagy. NFE2L2 is activated by reactive oxygen species-MAPK14 axis-dependent TLR4 signalling, and induces the accumulation of SQSTM1 and aggresome- like induced structures.
/ 5805	MI DZDZ	21220332	FLI1 contributes to lipopolysaccharide (LPS)-induced expression of matrix metalloprotease 1 (MMP-1), MMP-3, MMP-10, and interleukin-10 (IL-10) and rapid down-regulation of FLI1 expression after LPS stimulation attenuates the induction of various MMPs
75950	FLI1	20879862	and IL-10 under inflammatory conditions. PTCH1 is a member of the Hedgehog signalling pathway and is important for LPS-induced inflammatory response in macrophages. (Demonstrated in Murine model)
77092	PTCH1	21131441	STAT4 is a transcription factor that mediates IL12 cytokine signalling, and as a member of the STAT protein family, STAT4 have significant impact on
77625	STAT4	17971840	innate immunity during sepsis. STAT4 relies on the phosphorylation of it's N-domains to form STAT4 tetramer in response to cytokine-
77625	STAT4	14704793	induced activation. CTLA4 blockade abrogates protection by regulatory T cells in a mouse model of microbe-induced innate
79215	CTLA4	18824539	immune-driven colitis. CTNNAL1 is an alpha-catenin-related protein that shares structural similarities with cytoskeleton linker proteins that augments NF-kappaB activity, promotes
80038	CTNNAL1	17952117	cell migration and increases resistance to apoptosis.
IL8RB is a G protein-coupled receptor that mediates NADPH oxidase-independent neutrophil extracellular trap (NET) formation and this G protein-coupled receptor (GPCR) pathway is operative and drugsensitive in cystic fibrosis lung disease.

80804	IL8RB	20818377	sensitive in cystic fibrosis lung disease.
			AAMP over-expression inhibits NOD1 and NOD2
80875	AAMP	19535145	mediated NF-kappaB signalling
			NFATC2 interacts with JUN to synergistically activate
			interleukin-2 (IL2) transcription in T cells, a cytokine
			that acts as an autocrine growth factor during an immune response to bacterial and viral infection, as
81223	NFATC2	20557936	well as tumourigenesis.
01225	1011102	20337730	COPS8 is part of the COP9 signallosome that
84173	COPS8	20074051	functions to control NF-kappaB activation.
			LRRFIP1 interacts with and activates beta-catenin,
			which increases IFN-beta expression by binding to the
			C-terminal domain of the transcription factor IRF3 and
			recruiting the acetyltransferase EP300 to the IFN-beta
84322	LRRFIP1	20453844	enhanceosome via IRF3.
			LRRFIP1 (GCF2) acts as a repressor and occupies the -
			308 site of the TNF-alpha promoter in cells that do not make TNF-alpha. Other proteins may bind to the
			promoter, particular to the -308 site to transition from
84322	LRRFIP1	16199883	repressed to active transcription.
			LRRFIP1 is a regulator of toll-like receptor (TLR)
			pathway signaling and it co-localized with dsRNA in
84322	LRRFIP1	21102652	monocyte lysosomal structures.
			NFKB2 (p49/p100 subunit) associates efficiently with
07000		5025201	RELB and up-regulates the synthesis of NFKBIA
87893	NFKB2	7925301	(IKappaB-alpha).
			NFKB2 plays a key role in the regulation of RELA activation, suggesting an overlap in the function of NF-
			kappaB members in canonical and non-canonical
87893	NFKB2	18025196	pathway signalling.
			NFKB2 rearrangement gene product (p58) localizes in
			the nucleus to form a complex with RELA or RELB,
			suggesting that such NFKB2 gene rearrangement may
			therefore be a factor in the constitutive activation of
07000		10055400	NF-kappaB in adult T-cell leukemia (ATL), and
87893	NFKB2	18377428	thereby playing a role in the ATL pathogenesis. NFKB2 negatively regulates TCR signaling by binding
			with RELA, RELB, REL and NFKB1 (p50) in the
			cytoplasm and inhibiting these proteins from entering
87893	NFKB2	17548614	nucleus.
			NFKB2 limits TNF-induced bone resorption in mice
87893	NFKB2	19770515	by a TRAF3-dependent mechanism.
			VEGFA-induced tissue response, e.g. angiogenesis, is
			inhibited during RSV and Influenza viral infections.
			These effects were mediated by RIG-I and IFNR-
			dependent pathways, and consequently inhibited the Th2 inflammation response.
88716	VEGFA	21278304	m2 mnammanon response.
00/10	, 20171	212/050 F	

89810	NUP214	15210729	NUP214 and NUP153 are nucleoporins that control the nucleo-cytoplasmic shuttling and, along with XPO1- dependent nuclear export, the subcellular distribution of latent STAT1. MFN2 deletion causes inability of the cell to undergo mitochondrial fusion, and therefore exhibit impaired induction of interferons and pro-inflammatory cytokines in response to viral infection - which results
89999	MFN2	21285412	in increased viral replication. TNFRSF1B signalling induces selective BIRC2-
90091	TNFRSF1B	17220297	dependent ASK1 ubiquitination and terminates mitogen-activated protein kinase signalling.
90517	IL17F	19144317	IL17A and IL17F are required for the induction of beta-defensin in vivo in mice. IL17F signalling inhibits expression of pro-
90517	IL17F	19244213	<ul><li>inflammatory genes though sequential phosphorylation of CEBPB regulatory 2 domain.</li><li>IL17F is a Th17-related cytokine, traditionally thought of as an adaptive responder, has been shown to have</li></ul>
90517	IL17F	21074482	various innate sources and functions as a rapidly produced pro-inflammatory mediator. Innate IL17F- producing cells also employ many of the cytokine and transcriptional regulators utilized by Th17 cells. GRK5 is a serine/threonine kinase has a role in the regulation of G-protein coupled receptor (GPCR) signalling and is also an important regulator of signalling pathways stimulated by non-GPCRs. GRK5 also is a positive regulator of TLR4-induced I-kappa-B-
91131	GRK5	20945396	alpha-NFkappaB pathway as well as a key modulator of lipopolysaccharide-induced inflammatory response. DMBT1 encodes alternatively spliced proteins involved in mucosal innate immunity and two other molecules, a glycoprotein with a molecular mass of 340 kDa (GP340) and salivary agglutinin
91946	DMBT1	20418254	(DMBT1(SAG)). GP340 is secreted into broncho- alveolar surface lining fluid whereas DMBT1(SAG) is present in the saliva. Both interact with and agglutinate several Gram-negative and Gram-positive bacteria, as well as some viruses. GPSM1, an inhibitor of GNAI3, plays a key role in regulating autophagy; the dynamic interplay between GNAI3, GPSM1, and CCDC88A determines whether autophagy is promoted or inhibited. During starving conditions, GPSM1 binds to GNAI3 in MAP1LC3A- positive membranes to promote autophagy. When stimulated by growth factors, CCDC88A promotes the dissociation of GPSM1::GNAI3 complex to initiate anti-autophagy signalling pathways.
92256	GPSM1	21209316	anti-autophagy signalling pathways.

			CARD9 signalling mediates mammalian innate immune responses against selected fungi, bacteria, and viruses and can prime and shape adaptive immunity. CARD9 plays an essential role in downstream
92266	CARD9	19076343	signalling of the antifungal pattern-recognition receptor CLEC7A (Dectin-1). CARD9 is critical for full activation of innate immunity by converging signals downstream of multiple pattern recognition receptors (PRRs) and
92266	CARD9	20351059	plays a pivotal role in autonomous innate host defence against tuberculosis. FBXW5, an F-box family protein, negatively regulates
92843	FBXW5	19232515	MAP3K7 (TAK1) in the IL1B (IL-1beta) signalling pathway. IRAK1BP1 down-regulates Toll-like receptor-mediated transcription of several pro-inflammatory cytokines by changing the transcriptional profile of activated cells, leading to an increase in IL-10 production and
93345	IRAK1BP1	20534545	promoting LPS tolerance. Lysine-specific demethylase 1 (KDM1) is part of a repressor complex, along with key components that include histone deacetylase 1 or 2, RE-1 silencing transcription factor (REST), and co-repressor of REST (RCOR1). The KDM1/HDAC/REST/RCOR1
93808	KDM1	20798038	repressor complex is a significant component of host innate immunity. RUNX3 is capable of activating the CD11a gene promoter that directs CD11a/CD18 integrin expression as well as trans-activating the CD49d gene promoter. The leukocyte integrins CD11a/CD18 (LFA-1, alphaLbeta2) and CD49d (VLA-4, alpha4beta1, alpha4beta7) mediate leukocyte transendothelial migration during immune and inflammatory responses
94257	RUNX3	16164020	and provide co-stimulatory signals for the activation of T lymphocytes. HDAC1 Inhibition decreases IFN-alpha responsiveness whereas its expression augments the IFN-alpha
95712	HDAC1	14645718	response, demonstrating that it modulates IFN-alpha- induced transcription. Histone deacetylase 1 (HDAC1) is part of a repressor complex, along with key components that include HDAC2, RE-1 silencing transcription factor (REST), co-repressor of REST (CoREST), and lysine-specific demethylase (LSD) 1. The HDAC/CoREST/REST/LSD1 repressor complex is a
95712	HDAC1	20798038	significant component of host innate immunity. GJA1 plays an important role in innate immune control of commensal-mediated intestinal epithelial wound
95984	GJA1	19528242	repair.

			MAP3K7IP2 (TAB2) interacts with both MAP3K7 (TAK1) and TRAF6 and promotes their association, thereby triggering subsequent interleukin-1 signalling
97690	MAP3K7IP2	11259596	events. MAP3K7IP2 (TAB2) and MAP3K7IP2 (TAB3)
97690	MAP3K7IP2	15327770	activate the NF-kappaB pathway through binding to K63-linked polyubiquitin chains. MAP3K7IP2 binds to K-63 polyubiquitinated TRAF2
97690	MAP3K7IP2	19150425	and this association is required for activation of downstream IKK and JNK kinases. MAP3K7IP2 (TAB2) activates MAP3K7 (TAK1) and also plays an essential role in the deactivation of TAK1
97690	MAP3K7IP2	19955178	by recruiting PP6 through a polyubiquitin chain- dependent mechanism. MAP3K7IP2 is an adapter linking MAP3K7 (TAK1) and TRAF6 and also functions as a mediator of TAK1
97690	MAP3K7IP2	10882101	activation in the IL-1 signalling pathway. MAP3K7IP2 (TAB2) and MAP3K7IP3 (TAB3)
97690	MAP3K7IP2	14633987	function redundantly as mediators of TAK1 activation in IL-1 and TNF signal transduction. MKNK1 plays an important role in IFNG induced
98137	MKNK1	21149447	IRF1 expression and is essential for generation of anti- proliferative responses. MAP3K4 (MEKK4) dimerization is regulated both positively and negatively by its interaction with specific proteins. GSK3B negatively regulates MAP3K4 stimulation of MAPK14 (P38) and MAPK8
98770	MAP3K4	17726008	(JNK) activity. MAP3K4 sequesters RIPK2 to inhibit the
98770	MAP3K4	18775659	NOD2:RIPK2 complex from activating NF-kappaB signalling pathways. JUN interacts with NFATC2 to synergistically activate interleukin-2 (IL2) transcription in T cells, a cytokine that acts as an autocrine growth factor during an immune response to bacterial and viral infection, as
99221	JUN	20557936	well as tumourigenesis. GNAI3 is a G protein involved in autophagy signalling pathways and is tightly regulated through its interaction with its GEF activator (CCDC88A) and GDI inhibitor (GPSM1). During starving conditions, GPSM1 binds to GNAI3 in MAP1LC3A-positive membranes to promote autophagy. When stimulated by growth factors, CCDC88A promotes the dissociation of GPSM1::GNAI3 complex to initiate anti-autophagy signalling pathways.
100849	GNAI3	21209316	<u>o</u> 6 Fann

Gnai3 is a G protein involved in anti-autophagy signalling pathways and is tightly regulated through its interaction with its GEF activator (Ccdc88a) and GDI inhibitor (Gpsm1). During starving conditions, Gpsm1 binds to Gnai3 in Map1lc3a-positive membranes to promote autophagy. When stimulated by growth factors, Ccdc88a promotes the dissociation of Gpsm1::Gnai3 complex to initiate anti-autophagy signalling pathways.

10(205	с ·з	21200216	signannig painways.
186305	Gnai3	21209316	
101004	CD53	20407468	CD53 is an important regulator of innate tumour necrosis factor (TNF)-alpha levels. SNX27 co-localizes with CYTIP at the early endosomal compartment of lymphocytes suggesting a
			role for this interaction in endocytic trafficking and/or
102394	SNX27	17577583	signalling RUSC1 binds to IKBKG and TRAF6, representing a
			molecular link between NGF signalling and IKK
103247	RUSC1	19365808	complex activity.
			FCGR2A is a receptor for immunoglobulin G,
			clustering induces shape change, secretion and
104282	FCGR2A	12857726	aggregation
			FCGR2A (FcgammaRIIA) is a receptor that recognizes
			IgG opsonized particles and initiates phagocytosis in
			immune clearance. FCGR2A is also a positive
104282	FCGR2A	21044955	regulator of complement-mediated phagocytosis.
105504	CEU	10100450	CFH is a major regulatory protein that down-regulates
105504	CFH	18190458	alternative complement activation.
			CFH, a complement regulatory factor, interacts with
			host cell surfaces as well as C3d part of C3b and plays a major in the distinguishing host from non-host
			surfaces during the alternative activated complement
			pathway.
105504	CFH	21285368	puttivuy.
100001	UT II	21202200	PTPRC (CD45) suppresses JAK (Janus kinase) kinases
			and negatively regulates cytokine receptor signaling by
			regulating interleukin-3-mediated cellular proliferation,
			erythropoietin-dependent haematopoiesis and antiviral
105609	PTPRC	11201744	responses in vitro and in vivo.
			TRAF5 is a signal transducer for the TNF receptor
			superfamily that is involved in TNF-induced NF-
106494	TRAF5	11479302	kappaB activation and protection from cell death.
			TRAF5, along with other TRAFs, is recruited to many
			TNF-receptor (TNF-R) superfamily members and is an important modulator of the proving signaling events
			important modulator of the proximal signaling events that occur at the time of receptor engagement and
			activation. TRAF5 has been shown to be a positive
			regulator of a number of these receptors that are
106494	TRAF5	19017969	involved in T cell co-stimulation.

106494	TRAF5	20161788	TRAF5, a ubiquitin ligase, is a key molecule in the innate response against viral infection where it mediates the activation of IRF3 and NF-kappaB downstream of MAVS through the recruitment of NEMO.
			Hsa-mir-146b targets IRAK1 and TRAF6 mRNA and their expression is knocked down by co-expression of
127623	hsa-mir-146b	16885212	here expression is knocked down by co expression of hsa-mir-146b, showing that hsa-mir-146b can post- transcriptionally repress IRAK1 and TRAF6. MFN1 deletion causes inability of the cell to undergo mitochondrial fusion, and therefore exhibit impaired induction of interferons and pro-inflammatory cytokines in response to viral infection - which results in increased viral replication.
66114	MFN1	21285412	Mfn1 deletion causes inability of the cell to undergo
			mitochondrial fusion, and therefore exhibit impaired induction of interferons and pro-inflammatory cytokines in response to viral infection - which results in increased viral replication.
136722	Mfn1	21285412	
138525	Cd14	21078886	Cd14 contributes to nucleic acid uptake in macrophages and acts as a co-receptor for endosomal Tlr7/Tlr9 activation.
			Gpsm1, an inhibitor of Gnai3, plays a key role in regulating autophagy; the dynamic interplay between Gnai3, Gpsm1, and Ccdc88a determines whether autophagy is promoted or inhibited. During starving conditions, Gpsm1 binds to Gnai3 in Map1lc3a- positive membranes to promote autophagy. When stimulated by growth factors, Ccdc88a promotes the dissociation of Gpsm1::Gnai3 complex to initiate anti- autophagy signaling pathways.
148192	Gpsm1	21209316	Ccdc88a, an activator of Gnai3, plays a key role in
			regulating autophagy; the dynamic interplay between Gnai3, Gpsm1 and Ccdc88a determines whether autophagy is promoted or inhibited. When stimulated by growth factors, Ccdc88a disrupts the Gnai3::Gpsm1 complex, subsequently enhancing anti-autophagy signaling pathways and inhibits autophagy by activating Gnai3.
157020	Ccdc88a	21209316	Mknk1 plays an important role in Ifng induced Irf1 expression and is essential for generation of anti-proliferation responses.
176847	Mknk1	21149447	promeration responses.

20222	SPP1	21136203	SPP1 exist in secreted and intracellular form; intracellular SPP1, or iSPP1, is involved in cytoskeleton rearrangement and signal transduction downstream of innate immunity receptors (e.g. TLR). iSPP1 may also function as an adaptor or scaffolding protein.
29322	SPPI	21130205	Spp1 exist in secreted and intracellular form; intracellular Spp1, or iSpp1, is involved in cytoskeleton rearrangement and signal transduction downstream of innate immunity receptors (e.g. TLR). iSPP1 may also function as an adaptor or scaffolding protein.
185485	Spp1	21136203	Vegfa-induced tissue response, e.g. angiogenesis, is inhibited during RSV and Influenza viral infections. These effects were mediated by RIG-I and IFNR- dependent pathways, and consequently inhibited the Th2 inflammation response.
185847	Vegfa	21278304	C4bp plays an inhibitory role in antimicrobial response by delaying the classical complement activation and attenuating the lectin pathway activation. The major inhibitory role of C4bp is to facilitate the decay of C3 convertase.
190275	C4bp	21283780	C4BPB plays an inhibitory role in antimicrobial response by delaying the classical complement activation and attenuating the lectin pathway activation. The major inhibitory role of C4BPB is to facilitate the decay of C3 convertase.
106321	C4BPB	21283780	C4BPA plays an inhibitory role in antimicrobial response by delaying the classical complement activation and attenuating the lectin pathway activation. The major inhibitory role of C4BPA is to facilitate the decay of C3 convertase.
106328	C4BPA	21283780	APCS is a key negative regulator for innate immune responses to DNA and may be partly responsible for the insufficient immune responses after DNA vaccination in humans. APCS-DNA complex showed significant defects in innate immune activation, and specifically inhibits the functions of HMGB1 and
103905	APCS	21278351	antimicrobial peptide LL37. Kitl is a PI3K-activating ligand that increases the secretion of Il6 and Tnfa in LPS-stimulated mast cells, as well as the attenuating the production of Il1b.
187862	Kitl	21262348	

50398	KITLG	21262348	KITLG is a PI3K-activating ligand that increases the secretion of IL6 and TNFA in LPS-stimulated mast cells, as well as the attenuating the production of IL1B. (Demonstrated in murine model)
	-		Pdcd1 expression in T cells is induced by Ifna stimulation and provides feedback inhibition of T cell activation. It is proposed that strong innate inflammatory response (i.e. Ifna secretion) cause an attenuated T cell response by Pdcd1 in sustained immune reaction.
184148	Pdcd1	21263073	Pdcd1 is a negative immunomodulator that is upregulated by Hepatitis C virus to deliver negative signaling to Tlr-mediated pathways controlling expression of Il12, a key cytokine linking innate and adaptive immunity.
184148	Pdcd1	21263070	PDCD1 mediates functional impairment of early immune responses during HCV infection by limiting STAT1 phosphorylation, which consequently inhibits IL12 expression in monocytes / macrophages.
85346	PDCD1	21091911	PDCD1 expression in T cells is induced by IFNA stimulation and provides feedback inhibition of T cell activation. It is proposed that strong innate inflammatory response (i.e. IFNA secretion) cause an attenuated T cell response by PDCD1 in sustained immune reaction.
85346	PDCD1	21263073	PDCD1 is a negative immunomodulator that is upregulated by Hepatitis C virus to deliver negative signaling to TLR-mediated pathways controlling expression of IL12, a key cytokine linking innate and adaptive immunity.
85346	PDCD1	21263070	Rftn1 cooperates with the uptake receptor to mediate cell entry of poly(I:C), which is critical for activation of Tlr3 and the subsequent production of IFN and inflammatory cytokines.
190364	Rftn1	21266579	RFTN1 cooperates with the uptake receptor to mediate cell entry of poly(I:C), which is critical for activation of TLR3 and the subsequent production of IFN and inflammatory cytokines.
21351	RFTN1	21266579	Hspbp1 interacts with Pglyrp1 and inhibits the cytotoxic activity of the Hspa1a :: Pglyrp1 complex secreted by lymphocytes.
138595	Hspbp1	21247889	

70495	HSPBP1	21247889	HSPBP1 interacts with PGLYRP1 and inhibits the cytotoxic activity of the HSPA1A :: PGLYRP1 complex secreted by lymphocytes.
			Akap10 is required to induce Ptges2 in the synthesis of NO upon LPS stimulation, in addition Akap10 also mediates the Ptges2-induced expression of cytokines Il10 and Il6 in alveolar macrophages. (Demonstrated in rat models)
183923	Akap10	21247892	AKAP10 is required to induce PTGES2 in the synthesis of NO upon LPS stimulation, in addition AKAP10 also mediates the PTGES2-induced expression of cytokines IL10 and IL6 in alveolar macrophages. (Demonstrated in rat models)
35162	AKAP10	21247892	Ptges2 is responsible for nearly half of the increment in NO production by alveolar macrophages in response to LPS stimulation. The enhancing effect of Ptges2 on NO production is mediated through the ligation of Ptger2 and acting via PKA to induce cAMP production. In addition Ptges2 induces expression of cytokines Il10 and Il6, while inhibiting Tnfa. (Demonstrated in rat models)
160124	Ptges2	21247892	PTGES2 is responsible for nearly half of the increment in NO production by alveolar macrophages in response to LPS stimulation. The enhancing effect of PTGES2 on NO production is mediated through the ligation of PTGER2 and acting via PKA to induce cAMP production. In addition PTGES2 induces expression of cytokines IL10 and IL6, while inhibiting TNFA. (Demonstrated in rat models)
87283	PTGES2	21247892	Stim1 supports the influx of extracellular Ca(2+), which is required for production of reactive oxygen species in phagocytosis.
202769	Stim1	21239714	STIM1 supports the influx of extracellular Ca(2+), which is required for production of reactive oxygen species in phagocytosis.
24705	STIM1	21239714	Orail supports the influx of extracellular Ca(2+), which is required for production of reactive oxygen species in phagocytosis.
199057	Orai1	21239714	ORAI1 supports the influx of extracellular Ca(2+), which is required for production of reactive oxygen species in phagocytosis.
61853	ORAI1	21239714	-F H

			Itpr3 facilitates depletion of intracellular Ca(2+) for the internalization phase of FcgR-mediated phagocytosis.
156294	Itpr3	21239714	ITPR3 facilitates depletion of intracellular Ca(2+) for the internalization phase of FcgR-mediated phagocytosis.
82972	ITPR3	21239714	Itpr1 facilitates depletion of intracellular Ca(2+) for the internalization phase of FcgR-mediated phagocytosis.
174302	Itpr1	21239714	ITPR1 facilitates depletion of intracellular Ca(2+) for the internalization phase of FcgR-mediated
14187	ITPR1	21239714	phagocytosis. Bcl3 is critically involved in lung defense against
			Gram-negative bacteria Klebsiella pneumoniae, modulating functions of several cells to facilitate efficient clearance of bacteria. Loss of Bcl3 incurred dramatic cytokine imbalance in the lungs, failure to clear bacteria and increased susceptibility to K. pneumoniae pneumonia.
152576	Bcl3	21228348	Lrrfip2, upon phosphorylation, interacts with Myd88 during LPS stimulation and induces NFKB activity.
202692	Lrrfip2	21220426	LRRFIP2, when phosphorylated at serine 202, interacts with MYD88 during LPS stimulation to induces NFKB activity.
24724	LRRFIP2	21220426	TRIB3 is an inhibitor of TLR2 mediated NFKB activation and chemokine induction. Helicobacter pylori LPS stimulation decrease expression of TRIB3, which may be an important mechanism during H. pylori-associated pathogenesis.
209676	Trib3	21220698	TRIB3 is an inhibitor of TLR2 mediated NFKB activation and chemokine induction. Helicobacter pylori LPS stimulation decrease expression of TRIB3, which may be an important mechanism during H. pylori-associated pathogenesis.
37170	TRIB3	21220698	Chga undergoes proteolytic cleavage to give rise to biologically active peptides, including catestatin. Catestatin is a neuroendocrine peptide with effects on human autonomic function and has recently been found to be a cutaneous antimicrobial peptide by inducing mast cell activation. In mast cells, catestatin plays a role in the migration, degranulation and release of leukotriene and prostaglandins; in addition, catestatin also induces production of pro-inflammatory cytokines to prime the innate immune response.
165906	Chga	21214543	

CHGA undergoes proteolytic cleavage to give rise to biologically active peptides, including catestatin. Catestatin is a neuroendocrine peptide with effects on human autonomic function and has recently been found to be a cutaneous antimicrobial peptide by inducing mast cell activation. In mast cells, catestatin plays a role in the migration, degranulation and release of leukotriene and prostaglandins; in addition, catestatin also induces production of pro-inflammatory cytokines to prime the innate immune response.

17248 CHGA 21214543 Siglech is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction. 188669 Siglech 21208791 Siglece is upregulated and phosphorylated following lipopolysaccharide stimulation in order to limit TLRdriven cytokine production and help maintain a healthy 177258 Siglece 19933851 cytokine balance following infection. Siglece is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction. 177258 21208791 Siglece Siglec5 is a member of the rapidly evolving CD33related siglec subfamily that are expressed on cells of the innate immune system. Siglec5 mediates endocytosis of anti-Siglec5 and sialoside ligands and Sigelc5 endocytosis is clathrin and dynamin independent, requires ADP ribosylation factor 6, and 17562860 176568 Siglec5 traffics to lysosomes. Siglec5 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction. 176568 Siglec5 21208791 Siglec1 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction. 206881 Siglec1 21208791 SIGLEC15 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction. 2836 SIGLEC15 21208791 SIGLEC11 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction.

63914

SIGLEC11

21208791

			SIGLEC9 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction.
65664	SIGLEC9	21208791	SIGLEC8 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction.
65955	SIGLEC8	21208791	SIGLEC7 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction.
65680	SIGLEC7	21208791	SIGLEC6 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid autoimmune destruction.
66055	SIGLEC6	21208791	SIGLEC5 is bound by group B Streptococcus (GBS) beta protein and this functions to impair human leukocyte phagocytosis, oxidative burst, and
66180	SIGLEC5	19596804	<ul><li>extracellular trap production, promoting bacterial survival.</li><li>SIGLEC5 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role in the attenuation of innate immunity to avoid</li></ul>
66180	SIGLEC5	21208791	autoimmune destruction. SIGLEC1 is a member of the SIGLEC protein family that recognize sialoside-based patterns and plays a role
47695	SIGLEC1	21208791	<ul><li>in the attenuation of innate immunity to avoid autoimmune destruction.</li><li>SIGLEC10 is a member of the SIGLEC protein family that recognize sialoside-based patterns and responds</li></ul>
65888	SIGLEC10	21208791	selectively to danger associated molecular patterns (DAMPs) to initiate limited innate response. Rgmb is highly expressed in macrophages and
140416	Rgmb	21187450	negatively regulates II6 expressed in macrophages and dependent manner via MAPK and ERK pathway. RGMB is highly expressed in macrophages and
35165	RGMB	21187450	negatively regulates IL6 expression in a BMP ligand- dependent manner via MAPK and ERK pathway. (Demonstrated in murine model)
55105	NOWID	2110/430	

173238	4432412L15Rik	21187075	Palm3 is a LPS inducible gene that functions as an adaptor protein for Tlr4 signaling. Palm3 interacts with Sigirr to negatively regulate TLR signaling.
			Oas3 is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas3 activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production.
196327	Oas3	21190483	Oas2 is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas2 activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production.
196257	Oas2	21190483	Oas1h is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas1h activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production.
196607	Oas1h	21190483	Oas1g is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded
196639	Oas1g	21190483	RNA. Upon recognition of dsRNA, the Oas1g activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production.
			Oas1f is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas1f activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway
196587	Oas1f	21190483	triggers the RIG-I pathway and induce Ifnb production. Oas1e is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas1e activates Record to alcove asRNA. The Oas1e activates
196409	Oas1e	21190483	Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production. Oas1d is a pathogen recognition receptor for the viral
			pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas1d activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production.
196726	Oas1d	21190483	Oas1c is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas1c activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway
196449	Oaslc	21190483	triggers the RIG-I pathway and induce Ifnb production.

			Oas1b is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas1b activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production.
196543	Oas1b	21190483	Oas1a is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the Oas1a activates Rnasel to cleave ssRNA. The Oas/Rnasel pathway triggers the RIG-I pathway and induce Ifnb production.
196676	Oas1a	21190483	OAS2 is a pathogen recognition receptor for the viral pathogen-associated molecular pattern, double-stranded RNA. Upon recognition of dsRNA, the OAS2 activates RNASEL to cleave ssRNA. The OAS/RNASEL pathway triggers the RIG-I pathway and induce IFNB production.
58392	OAS2	21190483	Ctss is an endosomal and lysosomal protease that is upregulated during various inflammatory disorders. Tlr2, 3, 4 ligand engagement increases the proteolytic activities of Ctss in macrophages.
172850	Ctss	21145045	Ctsb is an endosomal and lysosomal protease that is upregulated during various inflammatory disorders. Tlr2, 3, 4 ligand engagement increases the proteolytic activities of Ctsb in macrophages.
172314	Ctsb	21145045	CTSB is a member of the Cathepsins protein family which are key modulators of cell death and
7654	CTSB	18762176	inflammatory responses. CTSB is an endosomal and lysosomal protease that is upregulated during various inflammatory disorders. TLR2, 3, 4 ligand engagement increases the proteolytic activities of CTSB in macrophages without changing the mRNA expression or the endogenous inhibitors of CTSB. (Demonstrated in murine model)
7654	CTSB	21145045	Nlrc5 is dispensable for cytokine induction in virus and bacterial infections under physiologic conditions.
182126	Nlrc5	21148033	Lair1 play an inhibition role in the mechanisms controlling Ifna production by pDCs both in normal and pathological innate immune responses.
137073	Lair1	21151495	LAIR1 play an inhibition role in the mechanisms controlling IFNA production by pDCs both in normal
68573	LAIR1	21151495	and pathological innate immune responses.

			IL19 cytokine is a member of the interleukin 10 (IL10) family that includes IL20, IL22, IL24, and IL26 and
106273	IL19	21157117	<ul><li>has been shown to negatively regulate the innate immune system.</li><li>C8B is the beta subunit of complement factor 8 (C8)</li></ul>
99157	C8B	12220191	and is one of five components that interact to form the membrane attack complex (MAC). DEFA4 along with DEFA1 and DEFA3 are cationic
6052	DEFA4	19024344	alpha-defensin peptides that have multiple functions in the immune system. PIK3R1 (p85-alpha) is a subunit of phosphoinositide 3-
25037	PIK3R1	17827709	kinase (PI3K) and PI3K is activated upon Toll-like receptor (TLR) ligation. PIK3R1 interacts with SYK during Fc-gamma receptor
25037	PIK3R1	16921024	phagocytosis and endocytosis. NLRP10 oligomerizes and binds to PYCARD, inhibits PYCARD mediated NF-kappaB activation and
30412	NLRP10	15096476	apoptosis, as well as CASP1-mediated IL1B maturation. NEU1 desialylation of sialyl alpha-2,3-linked beta-
300862	NEU1	19796680	galactosyl residues of TLR4 is essential for receptor activation and cellular signaling. CARD16 is an intracellular regulator of CASP1 activation and plays a role in the regulation of IL1B
69692	CARD16	11536016	<ul><li>secretion and NF-kappaB activation during the pro- inflammatory cytokine response.</li><li>PTPN11 is a STAT5A phosphatase that specifically interacts with STAT5A in vivo in a tyrosine</li></ul>
58105	PTPN11	12615921	phosphorylation-dependent manner, leading to the down-regulation of active STAT5A. PTPN11 (SHP-2) phosphotyrosine phosphatase
58105	PTPN11	8995399	<ul><li>interacts with JAK tyrosine kinases to induce cytokine signal transduction.</li><li>DUSP16 mitogen activated phosphotase that negatively regulates MAPK activity and acts as a</li></ul>
20236	DUSP16	11489891	<ul><li>shuttle protein, determining the localization of MAPKs in the cytoplasm.</li><li>BCL10 and MALT1 are essential mediators of NF-kappaB activation in response to the triggering of a diverse array of transmembrane receptors, including</li></ul>
100044	BCL10	18806265	antigen receptors. BCL10 mediates LPS-induced activation of NF-
100044	BCL10	17540779	kappaB and IL8 in human intestinal epithelial cells. NFKBIE is part of the IkappaB family of proteins that regulates NF-kappaB-dependent transcription by inhibiting DNA binding and localizing these factors to the cell cytoplasm, specifically by sequestering REL and RELA in the cytoplasm and inhibiting nuclear
89543	NFKBIE	11152669	export. OTUD7B is a deubiquitinating enzyme that participate in the resolution of inflammatory responses by
102040	OTUD7B	18178551	suppressing NF-kappaB.

			CFB is a crucial catalytic component of the C3
79667	CFB	17921140	convertase enzyme that activates the alternative pathway of complement-mediated immunity. ISG20 is an IFN-induced 3'-5' exonuclease that is
28713	ISG20	15064795	strongly induced by viral double stranded RNA and is activated by RELA/NFKB1. IL2RG physically associates with JAK2, suggesting
74884	IL2RG	8041779	that regulation of JAK2 may be linked to IL2 induced signal transduction. MST1R regulates endotoxin-induced innate immune
35720	MST1R	18684919	responses by regulating the production of and response to IFNG (IFN-gamma). ADAM10 associates with tetraspanins and regulates
14155	ADAM10	18981120	cleavage of TNF (TNF-alpha) and EGF (epidermal growth factor). MAP3K3 in its unphosphorylated form can form a complex with unphosphorylated MAP3K7 (TAK1) where non-phosphorylated MAP3K7 interferes with
63432	MAP3K3	18206350	MAP3K3 phosphorylation and NF-kappaB reporter activity. NKIRAS1, in both GDP-bound and GTP-bound states,
22073	NKIRAS1	15024091	<ul><li>inhibits phosphorylation of NFKBIB by IKBKB and makes it resistant to degradation.</li><li>KLK1 is part of the kallikrein-kinin system (KKS) that</li></ul>
64886	KLK1	18577888	is involved in cancer and angiogenesis. KLK1 is involved in the enzymatic cascade known as
64886	KLK1	18725990	the contact system in the blood which leads to an inflammatory response.
15857	C1S	16177097	C1S associates with C1R and C1Q to form the first component (C1) of the classical complement pathway. C1S is the modular serine protease responsible for
15857	C1S	16177097	cleavage of C4 and C2, the protein substrates for the first component (C1). PPP4C negatively regulates lipopolysacharide (LPS)-
25572	PPP4C	18634786	induced and TRAF6-mediated NF-kappaB activation by inhibiting the ubiquitination of TRAF6. MAP3K1 and TRAF6 play a pivotal role in the
22000	MAP3K1	18984593	retinoic-acid-inducible gene-I (RIG-I)-like helicase antiviral pathway. PTGS2 (COX-2) is suppressed through inhibiting the NF-kappaB activation by LPS and this may be
105426	PTGS2	18955820	associated with the anti-inflammatory effects of L. casei on Raw264.7 cells.
52534	KPNA1	16298512	<ul><li>KPNA1 (IPOA5) regulates the import of STAT proteins into the nucleus through nuclear pores.</li><li>KPNA1 contains a nuclear localization signal binding</li></ul>
52534	KPNA1	12740372	site for STAT1 homodimers, STAT1-STAT2 heterodimers and influenza A virus nucleoprotein. C8G is the gamma subunit of complement factor 8
92880	C8G	11058761	(C8) and is one of five components that interact to form the membrane attack complex (MAC).

			LAT dephosphorylation in human platelets is integrin- mediated and involves PTPN1 protein tyrosine phosphatase and both PTPN1 activation and LAT dephosphorylation processes were involved in the
239499	LAT	12857726	control of irreversible platelet aggregation upon FcgammaRIIa stimulation. C1R is the protease that mediates activation of the classical complement C1 complex and this catalytic property is mediated by C1R C-terminal region,
15930	C1R	11445589	comprising two complement control protein (CCP) modules followed by a serine protease (SP) domain. LY9 (SLAMF3) is part of the SLAM and SAP gene families that control innate and adaptive immune
104095	LY9	18501771	responses. Ptch is a member of the Hedgehog signaling pathway and is important for LPS-induced inflammatory
161037	Ptch1	21131441	response in macrophages. CFP binds to early apoptotic T cells and initiates
61793	CFP	18579773	complement activation, leading to C3B opsonization and ingestion by phagocytic cells. ICAM1 is an inducible surface glycoprotein belonging to the immunoglobulin superfamily that is involved in
26804	ICAM1	10924857	a wide range of inflammatory and immune responses. ICAM1 binds to ITGAL/ITGB2 (CD11a/CD18) and ITGAM/ITGB2 (CD11b/CD18) expressed by
26804	ICAM1	3349522	leukocytes and promotes their adhesion and transendothelial migration. HLA-E is a non-classical major histocompatibility complex (MHC) class I molecule that is the least polymorphic of all the MHC class I molecules and acts as a ligand for receptors of both the innate and the
299069	HLA-E	19000151	adaptive immune systems. EGR1 is an immediate early gene that is up-regulated by a multitude of growth factors, cytokines and
47191	EGR1	15545275	<ul> <li>environmental stresses to regulate LPS-induced SOCS1 transcription.</li> <li>IL6ST (GP130) is a common cytokine receptor subunit that is preferentially bound by SOCS3 on its SHP2-binding site, suggesting that some of the negative regulatory roles previously attributed to the phosphatase SHP-2 might in fact be caused by the</li> </ul>
21516	IL6ST	10829066	action of SOCS3. IGF1R is able to mediate activation of STAT3 in vitro
31432	IGF1R	10747872	and in vivo and the JAKs are essential for this process of activation. ELP2 is a STAT3-interacting protein that regulates
2565	ELP2	10954736	cytokine signal transduction and may regulate the activation of STAT3. F2RL2 is able to signal autonomously to induce IL8
29585	F2RL2	18264801	release mediated by ERK1/2 phosphorylation, which contributes actively to inflammatory responses.

35882	F2RL3	11907122	F2RL3 modulates inflammation in the lung where its activation stimulates IL6, IL8, and PTGER2 (prostaglandin E2) release from human respiratory epithelial cells. CD27 is a member of the tumour necrosis factor receptor superfamily that activates NF-kappaB and MADKO (ateras activated metain binance hum bl
14014	CD27	9582383	MAPK9 (stress-activated protein kinase/c-Jun N- terminal kinase) via TRAF2, TRAF5 and MAP3K14 (NIK). HOXA9 plays a role in the innate immune response to bacterial infection as a modulator of NF-kappaB-
10490	HOXA9	18981407	dependent transcription. GSTP1 suppresses LPS (lipopolysaccharide)-induced excessive production of pro-inflammatory factors by
60718	GSTP1	18962899	<ul><li>inhibiting LPS-stimulated MAPKs (mitogen-activated protein kinases) as well as NF-kappaB activation.</li><li>EPOR, a member of the cytokine receptor superfamily, associates with the JAK2 protein tyrosine kinase upon erythropoietin stimulation, inducing tyrosine</li></ul>
29281	EPOR	8068943	phosphorylation of cellular substrates, including the EPOR, to transduce a growth signal. WDR34 is a MAP3K7 (TAK1)-associated inhibitor of
88472	WDR34	19521662	the IL-1R/TLR3/TLR4-induced NF-kappaB activation pathway. ACAP1 (CENTB1) selectively down-regulates NF- kappaB activation via NODs pathways, creating a feedback loop and suggesting a novel role of ACAP1
24566	ACAP1	17005562	in innate immune responses to bacteria and inflammatory responses. ERAP1 associates with RBMX and a heterogeneous nuclear ribonucleoprotein to regulate both the constitutive release of TNFR1 exosome-like vesicles
34697	ERAP1	18445477	and the inducible proteolytic cleavage of TNFR1 ectodomains. CXCL11 exert direct antimicrobial effects in vitro against Bacillus anthracis spore and bacilli in a receptor-independent manner and contributes to pulmonary innate immunity. (Demonstrated in murine model)
25238	CXCL11	21124994	
			Cxcl11 exert direct antimicrobial effects in vitro against Bacillus anthracis spore and bacilli in a receptor-independent manner and contributes to pulmonary innate immunity.
179944	Cxcl11	21124994	CXCL9 exert direct antimicrobial effects in vitro against Bacillus anthracis spore and bacilli in a receptor-independent manner and contributes to pulmonary innate immunity. (Demonstrated in murine model)
25111	CXCL9	21124994	modelj

			Cxcl9 exert direct antimicrobial effects in vitro against Bacillus anthracis spores and bacilli in a receptor- independent manner and contributes to pulmonary innate immunity.
179870	Cxcl9	21124994	AP3B1 (AP-3), a lysosome-related organelle trafficking and biogenesis protein, is required for the production of pro-inflammatory cytokines in
30467	AP3B1	21045126	plasmacytoid dendritic cells upon recognition of viral nucleic acids by endosomal TLR7 or TLR9. AP3B1 is crucial for the trafficking of TLR9 to specific endosomal compartments for the induction of type I interferon.
30467	AP3B1	21119105	interferon.
169560	S100a10	21115493	S100a10 :: Anxa2 is a key profibrinolytic complex that assembles plasminogen and tissue plasminogen activator, and promotes plasmin generation. As a negative feedback regulation, plasmin can induce disassociation of the heterotetramer and ubiquitin- mediated degradation of S100a10.
			S100A10 :: ANXA2 is a key profibrinolytic complex that assembles plasminogen and tissue plasminogen activator, and promotes plasmin generation. As a negative feedback regulation, plasmin can induce disassociation of the heterotetramer and ubiquitin-
102454	S100A10	21115493	mediated degradation of S100A10. Anxa2 :: S100a10 is a key profibrinolytic complex that assembles plasminogen and tissue plasminogen activator, and promotes plasmin generation. As a negative feedback regulation, plasmin can induce serine phosphorylation of Anxa2 and prevents the translocation to cell surface.
181662	Anxa2	21115493	
14711	ANXA2	19965653	ANXA2 is found in a heterotetrameric complex with S100A10, serving as a key extracellular binding partner for pathogens and host proteins alike and can also can be shed or secreted. ANXA2 tetramer can activate human and murine macrophages through TLR4 demonstrating an important role for ANXA2 in the detection of danger to the host, whether from injury or invasion.
			ANXA2 :: S100A10 is a key profibrinolytic complex that assembles plasminogen and tissue plasminogen activator, and promotes plasmin generation. As a negative feedback regulation, plasmin can induce serine phosphorylation of ANXA2 and prevents the translocation to cell surface.
14711	ANXA2	21115493	

54381	CEACAM1	18836450	CEACAM1 inhibits TLR2 induced pro-inflammatory immune responses by inducing PTPN6-(SHP-1)- mediated inhibition of the PI3K-NF-kappaB signal transduction pathway in pulmonary epithelial cells.
156030	Tlr11	19801549	Tlr11 expression in epithelial cells is regulated by epithelium-specific Ets factors, ELF3 (ESE-1), EHF (ESE-3), via its Ets element and by IRF8 through its IRF motif. Tlr11 is an intracellular receptor in the endoplasmic
156030	Tlr11	21097503	reticulum, and acts as an innate sensor for Toxoplasma protein profilin. IL13RA1 is a receptor for IL13. Upon ligand engagement, IL13RA1 activates the transcription factor
83434	IL13RA1	21097505	STAT6 to induce M2 expression profile in macrophages. Il23a is a LPS induced gene, and its expression in
			macrophages correlate with the severity of chronic intestinal inflammation. Il23a is transcriptionally inhibited by the binding of Irf1 to the ISRE element and serve as a homeostatic checkpoint in chronic
197114	Il23a	21097874	intestinal inflammation. Sema3a is a ligand for Plxna4, and the overexpression of Sema3a exacerbate cytokine storm caused by Tlr agonist and bacterial sepsis. The expression of Sema3a
135492	Sema3a	21098092	can be induced by Tlr engagement, and thus completes an autocrine loop. SEMA3A is a ligand for PLXNA4, and the overexpression of SEMA3A exacerbate cytokine storm caused by TLR agonist and bacterial sepsis. The
24546	SEMA3A	21098092	expression of SEMA3A can be induced by TLR engagement, and thus completes an autocrine loop. Plxna4 is a receptor protein and its expression in macrophages is required for optimal cytokine
134005	Plxna4	21098092	production upon Tlr stimulation. Plxna4 is a crucial component of sepsis-induced cytokine storm by activating Rac1, Mapk8 and Nfkb. PLXNA4 is a receptor protein and its expression in macrophages is required for optimal cytokine
281690	PLXNA4	21098092	production upon TLR stimulation. PLXNA4 is a crucial component of sepsis-induced cytokine storm by activating RAC1, MAPK8 and NFKB. CASP6 plays an important role in the activation of alveolar macrophages by neutrophils. The cleavage of
33873	CASP6	21098228	<ul><li>IRAK3 by CASP6 mediates the degradation of NFKBIA and the induction of TNF upon stimulation by bacterial products.</li><li>Defa20 is a mouse paneth cell alpha-defensins that has selective bactericidal activities against intestinal</li></ul>
138822	Defa20	21099205	microbiota and these activities are dependent on the disulfide bonds.

			PELI2 and other pellino isoforms are the E3 ubiquitin ligases that mediate the IL-1-stimulated formation of K63-pUb-IRAK1 in cells, which may contribute to the
7478	PELI2	17997719	activation of IKBKB and NF-kappaB, as well as other signalling pathways dependent on IRAK1 and IRAK4. Hsa-mir-126 antagonism suppresses the effector function of TH2 cells and the development of allergic
126853	hsa-mir-126	19843690	airways disease. FFAR2 binding of short chain fatty acids (SCFAs) provides a molecular link between diet, gastrointestinal
44260	FFAR2	19865172	bacterial metabolism, and immune and inflammatory responses. NR1H4 is an essential component of a network of
52872	NR1H4	19864602	nuclear receptors that regulate intestinal innate immunity and homeostasis. Hsa-mir-98 and hsa-let-7e confer cholangiocyte expression of CISH in response to microbial challenge,
126843	hsa-mir-98	19592657	a process that may be relevant to the regulation of TLR-mediated epithelial innate immune response. CISH is a negative regulator for inflammatory cytokine signalling that enhances NFKBIA (IKappaB-alpha)
37175	CISH	19592657	degradation and regulation and activates NF-kappaB in cholangiocytes in response to LPS stimulation. CISH expression in cholangiocytes is conferred by hsa- mir-98 and hsa-let-7e in response to microbial
37175	CISH	19592657	challenge, a process that may be relevant to the regulation of TLR-mediated epithelial innate immune response. Hsa-let-7e and hsa-mir-98 confer cholangiocyte expression of CISH in response to microbial challenge,
126519	hsa-let-7e	19592657	a process that may be relevant to the regulation of TLR-mediated epithelial innate immune response. HMGB3 functions as a universal sentinel for nucleic-
88858	HMGB3	19890330	acid-mediated innate immune responses.
44657	HMGB2	19890330	HMGB2 functions as a universal sentinel for nucleic- acid-mediated innate immune responses. SIVA1 diminishes NFkappaB and enhances JNK
22673	SIVA1	19584092	activity to favour apoptosis by physically interacting with MAP3K7 (TAK1) and XIAP.
22673	SIVA1	19392652	SIVA1 promotes K-48 polyubiquitination of TRAF2 and inhibits TCR-mediated activation of NF-kappaB.
7946	SNAP23	18692471	SNAP23 phosphorylation by IKBKB regulates mast cell degranulation and anaphylactic reactions. IRF2 is a transcriptional repressor that functions by competing with the transcriptional activator IRF1 and has also been shown to functions as a transcriptional activator for several genes. IRF2 transcriptional activity is regulated by sumoylation whereby covalent modification of IRF2 by SUMO1 regults in enhanced
46310	IRF2	18514056	modification of IRF2 by SUMO1 results in enhanced transcriptional repression activity on IRF1.

			PLK1 is a IKKgamma/NEMO-binding domain (gammaBD) kinase, which negatively regulates TNF-
20740	PLK1	18957422	induced IKK activation and cyclin D1 expression.
83153	TRAF1	19698991	TRAF1 is a positive regulator of the NF-kappaB alternative pathway.
			TRAF1 mediates both the CD30 signaling-dependent and independent NF-kappaB activation, which prevents
			lymphoma cells from spontaneous and induced
83153	TRAF1	19540595	apoptosis. TRAF1 phosphorylation by the ubiquitously expressed
			kinase PKN1 plays a critical role in the negative
83153	TRAF1	18429822	regulation of tonic activity of JNK and NF-kappaB signalling pathways.
			PKN1 is a ubiquitously expressed kinase that, when phosphorylated by TRAF1, plays a critical role in the
			negative regulation of tonic activity of JNK and NF-
33268	PKN1	18429822	kappaB signalling pathways. FXR1 is an essential component of a network of
66609	FXR1	19864602	nuclear receptors that regulate intestinal innate
00009	ΓΛΚΙ	19804002	immunity and homeostasis. ERBB2IP ccts as a negative regulator of the MDP-
24405	ERBB2IP	16203728	mediated activation of NF-kappaB by NOD2. ERBB2IP overexpression inhibits NOD2-dependent
24405	ERBB2IP	16203728	cytokine secretion in mouse embryonic fibroblasts.
			RBCK1 (LUBAC) complexes with RNF31 and is involved in the physiological regulation of the
27270		10126069	canonical NF-kappaB activation pathway through
37270	RBCK1	19136968	linear polyubiquitylation of IKBKG. RNF31 complexes with RBCK1 (LUBAC) and is
			involved in the physiological regulation of the canonical NF-kappaB activation pathway through
3579	RNF31	19136968	linear polyubiquitylation of IKBKG.
			PIAS1 and other PIAS proteins have been shown to function as E3-type small ubiquitin-like modifier
			(SUMO) ligases, and sumoylation is a modulatory mechanism for several transcription factors.
18826	PIAS1	12855578	incentation for several transcription factors.
			PIAS1 is a protein inhibitor of activated STAT1 and arginine methylation of PIAS1 is essential for the
			repressive function of PRMT1 in interferon (IFN)-
			dependent transcription as well as for the recruitment of PIAS1 to STAT1 target gene promoters in the late
18826	PIAS1	19136629	phase of the IFN response. WDR62 is recruited to stress granules and mediates a
46287	WDR62	19910486	non-classical MAPK8 (JNK) activation.
			DAB2IP functions as ARF6-GTPase activating protein to negatively regulate phosphatidylinositol
02507		10040740	4, 5-bisphosphate (PIP2)-dependent TLR4-TIRAP-
83597	DAB2IP	19948740	MyD88 signaling.

			YJEFN3 is a downstream effector of anti-bacterial function in intestinal epithelial cells that is required for NOD2-mediated NF-kappaB activation following NOD2 mediated meansities of besterial means of the statement of the sta
39983	YJEFN3	15753091	NOD2 mediated recognition of bacterial muramyl dipeptide (MDP). YJEFN3 (NDUFA13) interacts with viral interferon regulatory factor 1 of Kaposi's sarcoma-associated
39983	YJEFN3	12163600	herpesvirus and inhibits interferon/retinoic acid- induced cell death. YJEFN3 belongs to the family of genes associated with retinoid-interferon mortality and serves as an essential component of the oxidative phosphorylation system. YJEFN3 interacts with human herpesvirus 6B U95 protein and the resulting silencing of U95 expression reduces viral load and abrogates loss of mitochondrial
39983	YJEFN3	17928352	membrane potential.
39983	YJEFN3	17823279	YJEFN3 blocks SRC-induced gene expression through STAT3 and activation of cell adhesion molecules. RP5-1000E10.4 (SIKE) is a physiological suppressor of IKKepsilon and TBK1 and plays an inhibitory role in virus- and TLR3-triggered IRF3 but not NF-kappaB
101319	RP5-1000E10.4	16281057	activation pathways. Trim30 negatively regulates TLR mediated NF-kappaB activation by targeting Map3k7ip2 (TAB2) and
204202	Trim30	18345001	Map3k7ip3 (TAB3) for degradation. AZI2 participates in both the TLR3-mediated and the cytoplasmic DDX58 (RIG-I) dsRNA recognition
23171	AZI2	17142768	pathways in type-1 interferon (IFN) induction by binding to MAVS, DDX58 and IFIH1 (MDA5). MAP3K7IP3 (TAB3) and MAP3K7IP2 (TAB2) activate the NF-kappaB pathway through binding to
53967	MAP3K7IP3	15327770	K63-linked polyubiquitin chains. MAP3K7IP3 (TAB3) and MAP3K7IP2 (TAB2) phosphorylation is mediated by SAPK2a/p38alpha and this contributes to the SAPK2a/p38alpha-mediated
53967	MAP3K7IP3	14670075	feedback control of MAP3K7 (TAK1) activity that also involves the phosphorylation of MAP3K7IP1 (TAB1). MAP3K7IP3 (TAB3) and MAP3K7IP2 (TAB2)
53967	MAP3K7IP3	14633987	function redundantly as mediators of MAP3K7 (TAK1) activation in IL-1 and TNF signal transduction. GLRX (GRX-1), a degluationylation enzyme, activates
34038	GLRX	21078302	TRAF6 and is responsible for IL1R / TLR-dependent induction of NFkB pathway. APOBEC3G;APOBEC3F (A3G) is an innate restriction factor that inhibits human immunodeficiency virus type 1 (HIV 1) replication
7998	BEC3G;APOBI	21078663	immunodeficiency virus type 1 (HIV-1) replication. Hsa-mir-152, hsa-mir-148a, and hsa-mir-148b, are negative regulators of the innate response and Ag- presenting capacity of dendritic cells (DCs), which may
127645	hsa-mir-152	21068402	contribute to the immune homeostasis and immune regulation.

			Hsa-mir-148b, hsa-mir-148a, and hsa-mir-152, are negative regulators of the innate response and Ag- presenting capacity of dendritic cells (DCs), which may contribute to the immune homeostasis and immune
126249	hsa-mir-148b	21068402	regulation. Hsa-mir-148a, hsa-mir-148b, and hsa-mir-152, are negative regulators of the innate response and Ag- presenting capacity of dendritic cells (DCs), which may
126833	hsa-mir-148a	21068402	contribute to the immune homeostasis and immune regulation. TRAT1 (TRIM56) is an interferon-inducible E3 ubiquitin ligase that modulates TMEM173 (STING) to confer double-stranded DNA-mediated innate immune regnonses
49131	TRAT1	21074459	responses.
			A coagulation factor II (thrombin) receptor belonging to a family of G protein-coupled receptors, protease- activated receptors (PARs), and during inflammation, microorganisms as well as host immune cells release various proteases activating protease-activated
29671	F2R	17977790	receptors (PARs) Activation of F2R stimulates IL6, IL8, and PTGER2
29671	F2R	11907122	(prostaglandin E2) release from human respiratory epithelial cells F2R (PAR1) activates MAPK14 (p38) and MAPK3/1
29671	F2R	21029417	<ul><li>(ERK1/2) kinases to trigger production of innate immunity markers in oral kertainocytes.</li><li>Slc15a4, a peptide/histidine transporter in organelle trafficking, is required for the production of pro-</li></ul>
201520	Slc15a4	21045126	inflammatory cytokines in plasmacytoid dendritic cells upon recognition of viral nucleic acids by endosomal Tlr7 or Tlr9. FCGR1A (FcgammaRI) is a receptor that recognizes IgG opsonized particles and initiates phagocytosis in
101961	FCGR1A	21044955	immune clearance. FCGR1A is also a negative regulator of complement-mediated phagocytosis. UBD mediates NF-kappaB activation and may promote
236915	UBD	19959714	tubulointerstitial inflammation in chronic kidney diseases. CD300E functions as an activating receptor capable of
67366	CD300E	20039296	regulating the innate immune response in myeloid cells. PRKCA is a key component that controls MyD88-
65194	PRKCA	19950169	dependent cytokine gene expression in human and mouse but differentially regulates production of TICAM1 (TRIF)-dependent cytokines. CASP10 is an initiator caspase in the death receptor (DR)-dependent apoptotic pathway with multiple identified splice variants. The prodomain-only isoform
78470	CASP10	17822854	of CASP10, Caspase-10g, may play a regulatory role in the NF-kappaB pathways.

106452	IRF6	12219090	IRF6 belongs to a family of transcription factors that share a highly conserved helix-turn-helix DNA-binding domain and a less conserved protein-binding domain. IRF9 is a DNA-binding protein that is a member of the alpha-interferon-induced ISGF3 complex. Bipartite
236583	IRF9	9242679	complexes of STAT2:IRF9 and STAT2:STAT1 translocate to the nucleus and associate on DNA target sites as ISGF3. IRF9 functions to recruit RNA polymerase II to the promoter of interferon (IFN)-stimulated genes and this
236583	IRF9	15194680	function of IRF9 requires the activity of histone deacetylases. VASP is an important component of the cellular microfilament system that plays a major role in the regulation of serum response element (SRE)-dependent
57716	VASP	14679200	transcription. VASP activity is regulated by GMP- dependent protein kinase (G-kinase). IFITM1 is an antiviral restriction factor that mediates cellular resistance to influenza A H1N1 virus, West
16232	IFITM1	20064371	Nile virus, and dengue virus by inhibiting their early replication. IFITM2 is an antiviral restriction factor that mediates cellular resistance to influenza A H1N1 virus, West
16199	IFITM2	20064371	Nile virus, and dengue virus by inhibiting their early replication.
301516	C4B	2650988	C4B is one of two isotypes of the fourth component of complement.
301516	C4B	12440962	C4B assembles with C2A to form the C3/C5 convertase that goes on to cleave complement C5. RAD21 is part of the cohesin complex which is the cellular machinery involved in sister chromatid cohesion and that which requires access to the nucleosomal DNA to perform its function in
33417	RAD21	12198550	chromosome segregation. RASGEF1B is a guanine-nucleotide exchange factor
26684	RASGEF1B	20090772	(GEF), whose expression is induced in macrophages on stimulation with toll-like receptor (TLR) agonists. NOXA1 together with NADPH oxidase organizer 1 (NOXO1) are key regulatory subunits of the NADPH oxidase NOX1, the activity of which is regulated through MAP kinase (MAPK), protein kinase C
93355	NOXA1	20110267	<ul><li>(PKC), and protein kinase A (PKA)-dependent phosphorylation on Ser-282 and Ser-172 of NOXA1.</li><li>TRPV2 has fundamental importance in innate</li></ul>
32212	TRPV2	20118928	immunity by participating in macrophage particle binding and early phagocytosis. CYBA is an NADPH oxidase and CYBA-dependent
46334	СҮВА	20142487	reactive oxygen species (ROS) are key regulators of neutrophil chemotactic migration.

			CALCA can be produced by immune cells such as monocytes/macrophages following inflammatory stimulation and has a positive or negative reciprocal effect on the production of other pro- and anti- inflammatory mediators, playing both facilitating and
33289	CALCA	20141542	suppressing roles in immune and inflammatory responses as shown in mice. PPP3CA (calcineurin) is a serine/threonine phosphatase that is activated by calcium and calmodulin that promotes HIF1A expression by
31818	PPP3CA	17965024	<ul><li>dephosphorylating RACK1 and blocking RACK1</li><li>dimerization.</li><li>KIR3DL2 functions as a CpG oligodeoxynucleotide</li><li>(ODN) receptor at the cell surface, facilitating the</li><li>encounter of CpG ODN with TLR9 in early</li></ul>
247106	KIR3DL2	20147700	endosomes.
54466	OPTN	20174559	OPTN (optineurin) has a role in the inhibition of virus- triggered IFNB1 induction.
			IL25, a member of the IL17 cytokine family, promotes the accumulation of a lineage-negative multipotent progenitor (MPP) cell population in the gut-associated lymphoid tissue that promotes T(H)2 cytokine responses, presenting an innate immune pathway that
3218	IL25	20200520	promotes T(H)2 cytokine responses at mucosal sites. LGALS4 (Gal-4), expressed in the intestinal tract, recognizes and kills human blood group antigen- expressing Escherichia coli while failing to alter the viability of other E. coli strains or other
49542	LGALS4	20154696	Gram-negative or Gram-positive organisms. LGALS4 is part of the galectin family of proteins that have emerged as autonomous bacteria-killing agents, pointing to a principal role of these proteins in innate
49542	LGALS4	20208507	immunity. LGALS8 (Gal-8), expressed in the intestinal tract, recognizes and kills human blood group antigen- expressing Escherichia coli while failing to alter the
107605	LGALS8	20154696	viability of other E. coli strains or other Gram-negative or Gram-positive organisms. LGALS8 is part of the galectin family of proteins that have emerged as autonomous bacteria-killing agents, pointing to a principal role of these proteins in innets
107605	LGALS8	20208507	pointing to a principal role of these proteins in innate immunity. RAC1 cooperates with TLR2, MyD88, and PI3K in
8115	RAC1	20167866	lipoteichoic acid-induced cPLA2/COX-2-dependent airway inflammatory responses. IL31 is a pruritogenic cytokine in human mast cells and its secretion is induced by antimicrobial peptides
62191	IL31	20190140	human beta-defensins and cathelicidin LL-37.

			RSAD2 (viperin) is an antiviral protein whose expression is highly upregulated during viral infections via IFN-dependent and/or IFN-independent pathways and an IRF1 mediated type I IFN independent mechanism of enhanced RSAD2 expression provides a
26912	RSAD2	20308629	redundant mechanism to protect cells from viral infections. ATF3 is induced by lipopolysaccharide (LPS) and regulates TLR-stimulated inflammatory responses as
106549	ATF3	16688168	part of a negative-feedback loop. HERC5 positively regulates the innate antiviral
29776	HERC5	20308324	responses by sustaining IRF3 activation via a novel post-translational modification, ISGylation. FCN2, as well as ficolins FCN1 and FCN3, in serum are associated with MBL-associated serine protease (MASP) to form a complex and this complex binds to carbohydrates present on the surface of a variety of Gram-positive and Gram-negative bacteria through ficolin, initiating complement activation via the lectin
91819	FCN2	20375620	pathway. DUOX1 and DUOX2 localize to the apical plasma membrane of epithelial cells in major airways, salivary glands, and the gastrointestinal tract, and provide extracellular hydrogen peroxide to lactoperoxidase to produce antimicrobial hypothiocyanite ions.
10154	DUOX1	18511861	Expression of dual oxidases DUOX2 and DUOX1 is regulated by Th1 and Th2 cytokines in human airways. NOX4 is implicated in innate immunity since lipopolysaccharide (LPS) induces NOX4-dependent
67316	NOX4	18511861	reactive oxygen species (ROS) generation. Abcg1 regulates innate immunity in a tissue-selective manner and Abcg1(-/-) mice have an enhanced pulmonary host defense response driven predominantly
163644	Abcg1	20395559	by hematopoietic cells. The Lyn/PI3K module negatively regulates activation
22229	LYN	20385881	of murine macrophages while Inpp5d (SHIP-1) promotes it. Inpp5d promotes while the Lyn/PI3K module
178553	Inpp5d	20385881	negatively regulates activation in murine macrophages viaTlr2 and Tlr4 receptors. INPP5D (SHIP-1) is a critical negative regulator of
83441	INPP5D	20100929	IFN-beta production downstream of TLR3 through the regulation of TBK1 localization and activity. INPP5D is a negative regulator of GM-CSF-derived
83441	INPP5D	20154203	dendritic cell (DC) generation but a positive regulator of GM-CSF-derived DC maturation and function. CNPY3 differentially interacts with TLR2, TLR4, and TLR9 and a single-nucleotide change in the CNPY3 gene con influence the strength of TLP responses and
87405	CNPY3	18780723	gene can influence the strength of TLR responses and may also alter the relative activity of each TLR.

CNPY3 (PRAT4A) is required for TLR-dependent immune responses where it regulates the subcellular distribution and response of multiple TLRs, required for both innate and adaptive immune responses.

BIRC5 is cleaved by GZMM (granzyme M) and this triggers degradation of the BIRC5-XIAP complex to 20406824 free caspase activity, leading to cytolysis of target cells. DHCR24 (Seladin1) is a novel lipopolysaccharide (LPS)-responsive gene and inhibits the tumour necrosis factor-alpha production and osteoclast formation in response to LPS. DHCR24 is an LPS-responsible gene product that negatively regulates the LPS-induced 20406300 inflammatory response.

> Gpr33 is highly expressed in murine dendritic cells and its expression is regulated by the activity of toll-like receptors (TLR) and AP-1/NF-kappaB signaling pathways in cell culture and in vivo.

> GPR33 is an orphan member of the chemokine-like receptor family and is a pseudogene in most humans.

> SFTPA2 is a carbohydrate pattern recognition molecule of innate immunity, that significantly enhances phagocytosis and killing of Aspergillus fumigatus, a pathogenic fungus, by neutrophils and macrophages.

> IFNGR1, along with IFNGR2, are the receptor subunits for IFN-gamma that upon ligand binding, translocate to the nucleus together with STAT1-alpha, and associate with IFN-gamma-activated sequence (GAS) elements at the promoter sites of IFN-gammaactivated genes.

> MIF;SLC2A11 interacts with ribosomal protein S19 (RPS19) leading to attenuation of MIF proinflammatory function by inhibition of the MIF-CD74 interaction and MIF triggered adhesion of monocytes.

MIF;SLC2A11 is a regulator of innate immunity and inflammation by having an important role in proinflammatory macrophage responses. Transcription factors specificity protein 1 (SP1) and cAMP response element-binding protein (CREB1) are critical positive regulators of its constitutive gene expression.

ILF3 is an RNA-binding protein that influences mRNA turnover and/or translation by regulating mRNA stability. ILF3 can bind to mitogen-activated protein (MAP) kinase phosphatase 1 (MKP-1) and increase its mRNA stability and translation, resulting in increased dephosphorylation activity and thereby MKP-1 inactivation of MAP kinases extracellular signalregulated kinase (ERK), c-Jun N-terminal kinase 18490444 (JNK), and p38.

138962	Gpr33	20399748
231020	GPR33	20399748
80070	SFTPA2	20413160

CNPY3

BIRC5

DHCR24

17998391

87405

70554

99054

- 97020 IFNGR1 16785527
- - MIF;SLC2A11 19155217

18034423

2604

27975

2604

ILF3

MIF;SLC2A11

			ATG9A is an essential autophagy protein that functions as a regulator of innate immunity following double stranded DNA stimulation by controlling dsDNA- driven dynamic translocation of stimulator of IFN
81394	ATG9A	19926846	genes (STING), aka TMEM173, and TBK1. MX2 is part of the Mx GTPase family of protein that are interferon-induced members of the dynamin superfamily of large GTPases, which inhibit a wide
3922	MX2	18062906	range of viruses by blocking an early stage of the replication cycle. TPSB2 (tryptase) expression is modulated by IL33, a novel pathway by which mesenchymal cells exposed to
8619	TPSB2	20427273	inflammatory cytokines modulate the phenotype of local MCs to shape their immune responses. GAB1 inhibits vesicular stomatitis virus (VSV) replication and VSV infection-induced cell damage by inducing type I IFNs and IFN-inducible gene expression via the PI3K/Akt pathway. It is needed for full activation of TLR3/4- and RIG-I-triggered innate
39257	GAB1	20435932	responses by promoting activation of PI3K/Akt, MAPKs, and NF-kappaB pathways. SLAMF1 is one of nine SLAM-family genes, a subfamily of the immunoglobulin superfamily, that encode differentially expressed cell-surface receptors of hematopoietic cells. SLAM and SLAM-associated protein (SAP) gene families control innate and adaptive
104073	SLAMF1	18501771	immune responses SLAMF1 is one of nine SLAM-family genes and SLAM receptors and SLAM-associated proteins (SAPs) influence lymphocyte interactions,
104073	SLAMF1	19079134	development and function. GOPC (cystic fibrosis transmembrane regulator- associated ligand) interacts with human papillomavirus type 16 E6 protein to promote E6-associated protein (E6AP)-mediated ubiquitination and proteasomal
95826	GOPC	16878151	degradation. YWHAE (14-3-3 epsilon) interacts with key components of mitogen-activated protein kinase (MAPK)signal module for selective modulation of the TNF-alpha-induced time course-dependent NF-kappaB
14629	YWHAE	20462248	activity. LTB4R (BLT1) is a high-affinity leukotriene B4 (LTB4) receptor that is expressed in a variety of immune cells such as neutrophils, macrophages and
236774	LTB4R	20959460	dendritic cells. LTB4-LTB4R signaling plays a pivotal role in macrophage phagocytosis and innate immunity. BTN3A3 is part of the BT3 family of immunoreceptors belonging to the extended B7 family that are expressed on the surface of resting and activated monocytes and monocyte-derived dendritic cells (iDC). BT3 molecules
69364	BTN3A3	20947169	are involved in the regulation of the balance between immune activation and suppression.

69145	BTN3A2	20947169	BTN3A2 is part of the BT3 family of immunoreceptors belonging to the extended B7 family that are expressed on the surface of resting and activated monocytes and monocyte-derived dendritic cells (iDC). BT3 molecules are involved in the regulation of the balance between immune activation and suppression. BTN3A1 is part of the BT3 family of immunoreceptors belonging to the extended B7 family that are expressed on the surface of resting and activated monocytes and monocyte-derived dendritic cells (iDC). BT3 molecules
69277	BTN3A1	20947169	are involved in the regulation of the balance between immune activation and suppression.
94975	IFI6	15685448	<ul><li>IFI6 has a function as a cell survival protein by inhibiting mitochondrial-mediated apoptosis.</li><li>IFI6 (G1P3) gene encodes a low molecular weight mitochondrial protein that stabilizes mitochondrial</li></ul>
94975	IFI6	20939681	function and opposes apoptosis. PTMA interaction with STAT3 is IFN-induced and
83127	РТМА	15242774	results in the nuclear translocation of the complex. PTMA inhibits HIV-1 via Toll-like receptor 4- mediated type I interferon (IFN) induction by acting as a ligand for TLR4 and stimulating type I IFN production to potently suppress HIV-1 after entry into
83127	РТМА	20479248	cells. Rxra controls innate inflammatory responses through the up-regulation of chemokine expression. Mice lacking Rxra in myeloid cells exhibit reduced levels of CCL6 and CCL9, impaired recruitment of leukocytes to sites of inflammation, and lower susceptibility to
151388	Rxra	20498053	sepsis. RXRA controls innate inflammatory responses through the up-regulation of chemokine expression. Mice lacking RXRA in myeloid cells exhibit reduced levels of CCL6 and CCL9, impaired recruitment of leukocytes to sites of inflammation, and lower
91747	RXRA	20498053	susceptibility to sepsis. Nfatc2 is a cytokine that acts as an autocrine growth factor during an immune response to bacterial and viral infection, as well as tumorigenesis.
213130	Nfatc2	20557936	AIMP1 is phosphorylated by JNK through the TLR- MyD88 pathway, leading to the loss of its regulatory activity for endoplasmic reticulum retention of gp96 and resulting in the increase of cell surface expression of gp96, thus providing a new molecular mechanism
33233	AIMP1	20510162	underlying TLR-mediated gp96 regulation. GNB2 is the major G protein isoform that mediates neutrophils directional cell migration and in vivo
31798	GNB2	20525682	infiltration.

			SPI1 (PU.1) directly regulates FLT3 kinase in a concentration-dependent manner, and is a critical
			regulator of both conventional and plasmacytoid
43260	SPI1	20510871	dendritic cell development.
			NLRP7 is a member of the PYRIN-containing
			apoptotic protease-activating factor-1-like proteins that
(0020	NU DD7	15017402	functions as a feedback regulator of CASP1-dependent
69828	NLRP7	15817483	interleukin-1 beta secretion.
			GLI1 functions as an antagonist of NF-kB activity after LPS stimulation at the level of promoter binding. GLI1
			interacts with RELA (p65) upon LPS stimulation and
			inhibits RELA-mediated transcriptional transactivation
			by interfering with RELA binding to target gene
42577	GLI1	20547752	promoter DNA.
			MARCO is upregulated in TLR4-mediated LPS
			responses and these receptors contribute to the efficient
			capturing and clearance of invading microbial
67847	MARCO	20162551	pathogens.
			TOMM70A, a mitochondrial import receptor, interacts
17222	TOMM70A	20628368	with MAVS upon RNA virus infection acting as a critical adaptor bridging TBK1/IRF3 to MAVS.
47323	TOMIM/0A	20028308	USP17 is required for virus-induced RIG-I- and
			melanoma differentiation-associated protein-5
			(MDA5)-mediated type I IFN signaling and functions
			through deubiquitination of RIG-I and MDA5 to
298136	USP17	20368735	regulate virus-induced type I IFN signaling.
			Ace and iNOS overexpression by myelomonocytic
			cells substantially boosts innate immunity and
			represents a new means to address serious bacterial
213293	1 00	20937811	infections such as L. monocytogenes and methicillin resistant S. Aureus.
213293	Ace	20937811	ETS1 transcription factor blocks terminal
			differentiation of keratinocytes and induces expression
			of matrix metalloproteases and innate immune
75897	ETS1	20930145	mediators.
			RAB11A is an important regulator of Toll-like receptor
			4 (TLR4) and TRAM transport to E. coli phagosomes
			thereby controlling IRF3 activation from this
17737	RAB11A	20933442	compartment.
			FZD1 has a role in the reciprocal regulation of the Toll- like receptor (TLR)/nuclear factor-kappaB (NF-
			kappaB) and the Wnt/beta-catenin pathway after
			aerosol infection of mice with Mycobacterium
25942	FZD1	20667980	tuberculosis.

			Fzd1 has a role in the reciprocal regulation of the Toll- like receptor (TLR)/nuclear factor-kappaB (NF- kappaB) and the Wnt/beta-catenin pathway after aerosol infection of mice with Mycobacterium tuberculosis. Fzd1 mRNA was significantly up- regulated during the course of infection in mice and its induction was dependent on TLRs, the myeloid differentiation response gene 88 (MyD88), and a
132454	Fzd1	20667980	functional NF-kappaB pathway. CTSG, a cathepsin, is a key modulator of cell death and
4005	CTSG	18762176	inflammatory responses. ELANE and CTSG, together with externalized nucleosomes, promote coagulation and intravascular thrombus growth in vivo. During systemic infection, activation of coagulation fosters compartmentalization of bacteria in liver microvessels and reduces bacterial
4005	CTSG	20676107	invasion into tissue. ELANE is a pro-inflammatory protease that regulates IL8 production from airway epithelial cells and can activate both EGFR and TLR4.
13111	ELANE	18772136	
13111	ELANE	18802098	ELANE mediates innate host protection against Pseudomonas aeruginosa by degrading the major outer membrane protein F, a protein involved in porin activity, maintenance of structural integrity, and sensing of host immune system activation. ELANE and CTSG, together with externalized nucleosomes, promote coagulation and intravascular thrombus growth in vivo. During systemic infection, activation of coagulation fosters compartmentalization
13111	ELANE	20676107	of bacteria in liver microvessels and reduces bacterial invasion into tissue. TNFRSF13B (TACI) triggers class-switch recombination (CSR) via the DNA-editing enzyme AID by activating NF-kappaB through a Toll-like
32554	TNFRSF13B	20676093	receptor (TLR)-like MyD88-IRAK1-IRAK4-TRAF6- TAK1 pathway. DHX9 interacts with CpG-B and was associated with TNF-alpha and IL-6 production and NF-kappaB activation upon CpG-B stimulation. DHX9 is localized
105280	DHX9	20696886	in the cytosol and is found to bind to TLR domain of MYD88. MAP2K1 and MAP2K2 are necessary and sufficient for the direct hinding of the mitagen estimated matching.
18019	MAP2K1	11134045	for the direct binding of the mitogen-activated protein kinases (MAPKs) MAPK3 (ERK1) and MAPK1 (ERK2). DDIT3 (CHOP) is an endoplasmic reticulum (ER) stress-induced transcription factor that targets the IL23 gene and this binding is enhanced in the context of
42805	DDIT3	20876114	both ER stress and Toll-like receptor (TLR) stimulation.

			ATG16L1 is involved in autophagy where it is
83447	ATG16L1	19898471	recruited to the plasma membrane at the site of bacterial entry by NOD1 and NOD2. ATG16L1, a critical autophagy protein, is recruited to
83447	ATG16L1	20200479	the plasma membrane by NOD2 during bacterial invasion. OTUB2 and OTUB1 negatively regulate virus-
17807	OTUB2	19996094	triggered type I interferon (IFN) induction and cellular antiviral response by deubiquitinating TRAF3 and -6. OTUB1 and OTUB2 negatively regulate virus-
52992	OTUB1	19996094	triggered type I interferon (IFN) induction and cellular antiviral response by deubiquitinating TRAF3 and -6. Padi4 plays a role in chromatin decondensation to form
201304	Padi4	20733033	neutrophil extracellular traps (NETs) in an innate immune response to bacterial infection. IL21 is a cytokine that has broad effects on both innate
36977	IL21	20817119	and adaptive immune responses. PPIA (CYPA) is a cell-intrinsic sensor for human immunodeficiency virus 1 (HIV-1) that exists in dendritic cells and mediates an antiviral innate immune
15159	PPIA	20829794	response. CD200 is induced by TLR-, NOD2-, and NALP3-
49799	CD200	20833375	mediated pathways, limiting macrophage activation and protecting the host from excessive inflammation. ANXA1 is cleaved by CAPN1 to generate a N- terminally truncated form of ANXA1 shown to be anti- inflammatory and able to activate ERK. This C- terminal ANXA1 peptide functions by increasing ICAM1 clustering around adherent neutrophils to anchor them to the endothelium and promote
71054	ANXA1	20679535	transmigration through the transcellular route. XDH (xanthine oxidase) is involved in TLR7/8-
42778	XDH	20632067	mediated activation of CASP1 and IL1B in an HIF1A- dependent manner. CR2 (CD21) is a cell membrane receptor, with 15 or 16 extracellular short consensus repeats (SCRs), that promotes B lymphocyte responses and bridges innate
106349	CR2	20558730	and acquired immunity. The interferon alpha receptor is composed of two subunits: IFNAR1 and IFNAR2;IL10RB. Binding of interferon (IFN)-alpha to IFNAR2;IL10RB results in STAT2 binding and the initiation of the IFN signaling
1973	FNAR2;IL10R1	12220192	cascade. Type I interferons (IFNs) play an important role in innate immunity to protozoan parasites by binding the IFN alpha receptor, composed of IFNAR1 and IFNAR2;IL10RB, and regulating neutrophil/monocyte recruitment, neutrophil turnover, and Leishmania
1973	FNAR2;IL10RI	20483775	infection.

			Foxa2 is expressed selectively in the respiratory epithelium where it plays a critical role in regulating genetic programs influencing Th2 cell-mediated
208945	Foxa2	20483781	pulmonary inflammation. FOXA2 is expressed selectively in the respiratory epithelium where it plays a critical role in regulating
59234	FOXA2	20483781	genetic programs influencing Th2 cell-mediated pulmonary inflammation. Defb14 is a beta-defensin with direct antimicrobial properties that contribute to local innate immune responses and it aids in combating microbial invasion by being chemotactic for a broad spectrum of leukocytes in a CCR6- and CCR2-dependent manner.
137815	Defb14	20483750	Defb4 is a beta-defensin with direct antimicrobial properties that contribute to local innate immune responses and it aids in combating microbial invasion by being chemotactic for a broad spectrum of
137844	Defb4	20483750	leukocytes in a CCR6- and CCR2-dependent manner.
157011		20100700	Fcrl5 is an orphan immunoregulatory protein that is highly expressed by innate B lymphocytes, as a specific receptor for orthopoxvirus MHC class I-like protein (OMCP) and this strongly implicates it in contributing to host defense against zoonotic
157234	Fcrl5	20519648	orthopoxviruses. Ly86 (MD-1) complexes with Toll-like receptor homolog Cd180 (RP105) to regulate the Ly96 (MD- 2)/Tlr4-mediated lipopolysaccharide (LPS) response. Soluble Ly86 alone, in addition to its complex with
145270	Ly86	20534476	Cd180, can regulate host LPS sensitivity. ING4 is a member of the inhibitor of growth (ING) family of chromatin-modifying proteins that functions to negatively regulate the cytokine-mediated inflammatory response. In mice, it facilitates NF-
14716	ING4	20534538	<ul> <li>kappaB activation of IkappaB promoters, thereby suppressing nuclear RelA (p65) levels and the activation of select NF-kappaB target cytokines.</li> <li>Ing4 is a member of the inhibitor of growth (ING) family of chromatin-modifying proteins that functions to negatively regulate the cytokine-mediated inflammatory response in mice by facilitating NF-</li> </ul>
188872	Ing4	20534538	<ul><li>kappaB activation of IkappaB promoters; thereby suppressing nuclear Rela (p65) levels and the activation of select NF-kappaB target cytokines.</li><li>Calm1 (CaM) and Nos2 (iNOS) coordinately function to form a stable complex that is part of a rapid host response that functions within the first 30 min</li></ul>
163852	Calm1	16893173	following bacterial infection to upregulate the innate immune system involving macrophage activation.

CALM1 (CaM) and NOS2 (iNOS) coordinately function to form a stable complex that is part of a rapid host response that functions within the first 30 min following bacterial infection to upregulate the innate immune system involving macrophage activation.

15903	CALM1	16893173	immune system involving macrophage activation. NXN subfamily of proteins form a link between
14319	NXN	20400501	MYD88 and FLII (flightless I) to mediate negative regulation of the TLR4/MYD88 pathway. SPON2 expression is upregulated during intestinal
6462	SPON2	20205276	inflammation and may induce NF-kappaB promoter activation in a TLR-9 mediated manner. CTCF regulates the transcription of the interleukin 1
36980	CTCF	15670593	receptor-associates kinase 2 (IRAK2) promoter. IRAK2 is part of a family of four IRAKs that regulate immune responsiveness to bacterial endotoxins. RNF5 negatively regulates virus-triggered signaling by targeting TMEM173, an adaptor protein that links virus-sensing receptors to IRF3 activation, for
80442	RNF5	19285439	ubiquitination and degradation at the mitochondria. Functions as an E3 ligase to promote STAT1 SUMO
3013	PIAS2	12764129	modification
3611	TCF4	18854153	Transcription Factor that is an essential and specific regulator of Plasmacytoid dendritic cell development Tyrosine protein kinase that acts with AXL and
7206	TYRO3	18083102	MERTK as pleiotropic inhibitor of the innate immune response in DCs The FN14 cytoplasmic tail binds to tumour-necrosis-
11328	TNFRSF12A	12529173	factor-receptor-associated factors 1, 2, 3 and 5 and mediates nuclear factor-kappaB activation
13548	TLR10	15728506	Human TLR10 is an orphan member of the TLR family with no identified specific ligand NLRP3 has an important role in IL-1 beta and IL-18
			secretion through the inflammasome and mediates
13548	TLR10	18487086	responses to LPS, peptidoglycan, bacterial RNA and imidazoquinolines. Able to homodimerize, heterodimerize with TLR1 and
13548	TLR10	15728506	TLR2, and directly associate with MYD88 mmu-mir-155 suppresses Socs1 protein expression and
13548	TLR10	22043967	has a pro-inflammatory role in microglia. NLRP3 complexes with PYCARD (ASC), RIPK2, and
13548	TLR10	18280719	CASP1 inflammasome to process IL-1 beta. Hsa-mir-155 is induced by bacterial and viral infections
13548	TLR10	17911593	as well as pro-inflammatory cytokines and functions to suppress FADD, RIPK1 and IKKE expression. NLRP3 is a critical NOD-like receptor family member that transduces a fungal recognition signal to the
13548	TLR10	19339971	inflammasome adaptor PYCARD for CASP1 activation and pro-IL-1beta processing. Mmu-mir-155 has a pro-inflammatory role in
13548	TLR10	22170100	astrocytes and its expression is negatively regulated by Irf3. (Demonstrated in human)

NLRP3-dependent CASP1 activation complex (inflammasome) is triggered when dying tumor cells release ATP, acting on P2X7 purinergic receptors from dendritic cells and allowing for the secretion of IL-1beta.

13548	TLR10	19767732	
			Hsa-mir-155 induction in response to either poly(I : C) or TNF-alpha is blocked by pharmacological inhibition
12540	TI D 10	17242265	of JNK, suggesting that its inducing signals use the
13548	TLR10	17242365	JNK pathway. NALP3 inflammasome is activated when CyaA, a virulence factor from B. pertussis, promotes IL-1beta production, which then polarizes T cell responses toward the Th17 subtype and promotes clearance of the
13548	TLR10	20610650	bacteria from the respiratory tract. Mir155 antagonizes progesterone to reverse the
13548	TLR10	22546503	inhibition of Tlr3/4 signalling.
			NLRP3 is directly activated by certain antibiotics and plays an important role in the antibiotic-mediated secretion of IL1B. In the case of polymyxin B, NLRP3 was also required for the neutrophil influx into the peritoneal cavity. (Demonstrated in murine models)
13548	TLR10	21278344	perionear eavity. (Demonstrated in multice models)
			Hsa-mir-155 expression in monocyte and macrophage cell lines is simulated by viral or bacterial infection in
13548	TLR10	19008191	vitro. NLRP3 inflammasome is essential for host defence
13548	TLR10	21289120	against influenza and other RNA viruses (i.e. EMCV, VSV).
13548	TLR10	23572582	Mir155 targets Pmaip1 (Noxa) and Socs1 to mediate natural killer cell expansion during MCMV infection. NLRP3 recruits adaptor protein PYCARD and CASP1
13548	TLR10	21385879	to form an NLRP3 inflammasome complex in response to Varicella-Zoster Virus (VZV) infection. hsa-mir-155 feedback positively regulates host antiviral innate immune response by promoting type I IFN
13548	TLR10	20937844	signaling via targeting suppressor of cytokine signaling 1 (SOCS1). NLRP3 is a component of the inflammasome and is
13548	TLR10	21439959	required for inflammation in acute pancreatitis. (Demonstrated in murine model) Mycobacterial infection induces expression of Mir155,
			which promotes the maturation of phagosomes and represses the expression of Rheb by targeting its
13548	TLR10	24130493	3†UTR. NLRP3 is necessary to illicit IL1B response specific to viable, but not heat-killed, E. coli infections. (Demonstrated in murine model)
13548	TLR10	21602824	(Demonstrated in murine model)

12540		21097505	hsa-mir-155 target IL13RA1 and reduces the IL13RA1 protein expression, and inhibits the expression of M2/pro-Th2 profile genes in macrophages.
13548	TLR10	21097505	The NLRP3 inflammasome plays a role in innate immune responses against mucosal Candida infection. NLRP3 limits the severity of infection when present in
13548	TLR10	22174673	either the hematopoietic or stromal compartments. (Demonstrated in mouse) T-cell-intrinsic Mir155 is required for type-2 immunity, in part through regulation of S1pr1, whereas
13548	TLR10	25024218	T-cell-intrinsic Mir146 is required to prevent overt Th1/Th17 skewing. NLRP3/PYCARD inflammasome activation following human respiratory syncytial virus infection is dependent on the activation of TLR2/MYD88/NF-kB
13548	TLR10	22295065	and reactive oxygen species/potassium efflux. MIR155 upregulation is a feature of the mammalian inflammatory response and MIR155 expression may exert both positive and negative regulatory effects on TLR and NFkB signalling. MIR155 expression is
13548	TLR10	21652514	suppressed by IL10 as a part of negative-feedback loop in LPS-stimulated cells. MIR223 and EBV miR-BART15 regulate the NLRP3
13548	TLR10	22984081	inflammasome and IL-1beta production. hsa-mir-155 overexpression can enhance innate antiviral immunity by promoting the JAK/STAT signalling pathway to facilitate the clearance of
13548	TLR10	21762537	hepatitis B virus in human hepatoma cells. Uromodulin nanoparticles activate the NLRP3
13548	TLR10	22997256	inflammasome in renal interstitial monocytes. hsa-mir-155 suppresses SOCS1 protein expression and has a pro-inflammatory role in microglia.
13548	TLR10	22043967	(Demonstrated in mice) Protein-bound polysaccharide-K can activate the NLRP3 inflammasome and induce IL1B in a TLR2-
13548	TLR10	24323452	and NLRP3-dependent manner. MIR155 has a pro-inflammatory role in human astrocytes and its expression is negatively regulated by
13548	TLR10	22170100	IRF3. Endoplasmic reticulum (ER) stress-mediated reactive oxygen species accumulation leads to activation of
13548	TLR10	24217221	NLRP3 inflammasome through enhanced secretion of IL1B and binding of TXNIP. MIR155 antagonizes progesterone to reverse the inhibition of TLR3/4 signalling (Demonstrated in
13548	TLR10	22546503	inhibition of TLR3/4 signalling. (Demonstrated in mice) Macrophages sense multiple types of bacterially derived RNA (mRNA, tRNA and rRNA) via the
13548	TLR10	25355909	NLRP3 inflammasome.

			MIR155 exerts anti-HIV-1 effects by targeting several HIV-1 dependency factors involved in post-entry, pre-
13548	TLR10	23028330	integration events. NLRP3 mediates NF-kB activation in both sterile and
13548	TLR10	25761061	microbially induced inflammation. MIR155 is targeted by Borna disease virus (BDV)
13548	TLR10	23428672	encoded protein to inhibit type I IFN induction. Circulating MIR155 activates natural killer cells via the
13548	TLR10	23580661	TLR1 signalling pathway. Hepatitis C virus (HCV)-induced, MIR155-regulated
			HAVCR2 expression regulates natural killer cell function, suggesting a novel mechanism for balancing
13548	TLR10	25772938	immune clearance and immune injury during chronic viral infection.
			Mir155 is a post-transcriptional repressor of Arntl (Bmal1), linking the molecular clock and innate
13548	TLR10	25995365	immune response.
			Influenza A virus non-structural protein 1, NS1, physically interacts with endogenous NLRP3 downregulating NLRP3 inflammasome activation as well as NF-ΰB, leading to a reduction in the levels of
13548	TLR10	25978411	inflammatory cytokines.
			TLR10 is a functional receptor involved in the innate immune response to H. pylori infection and the
13548	TLR10	25977263	TLR2/TLR10 heterodimer functions in H. pylori lipopolysaccharide recognition. Cytoplasmic USP7 binds to and deubiquitinates
13958	USP7	18952891	TRAF6 and IKBKG, thus terminating TLR-mediated NF-kappaB and JNK activation
15750	0517	10/520/1	A microtubule-interacting protein that positively
18462	CARD6	16418290	regulates NF-kappaB activation and modulates function of RIP family members Expression of CARD6 and RIPK2 in bone marrow-
			derived macrophages is rapidly induced by IFNB1 and
18462	CARD6	18160713	IFNG Selectively modulates NF-kappaB activation by RIPK2
18462	CARD6	12775719	and NOD1 Complement component seven is one of five
18519	C7	11058761	components that interact to form the cytolytic membrane attack complex (MAC)
			Complement component six is one of five components that interact to form the cytolytic membrane attack
18604	C6	11058761	complex (MAC) Involved in the early immune response against Listeria
18990	BDKRB2	18810490	infection by increasing the production of IL12A/IL12B in human monocyte-derived dentritic cells
			Cathepsins: key modulators of cell death and
19613	CTSD	18762176	inflammatory responses

			Binds proteins involved in complement, coagulation, and kinin systems, as well as viral and bacterial pathogens including S. aureus protein A and when expressed on activated platelets may contribute directly to thrombosis, inflammation and endovascular
21497	C1QBP	12574814	infections
22594	CSK	15749833	Plays a critical role in IL-1-induced NF-kappaB activation through the IKK complex Promotes calcium-regulated exocytosis of signal peptide-containing cytokine secretion (CCL5 but not IL-1beta) in human monocytes and mouse
22989	SCAMP5	19234194	macrophages Activates MEFV by binding to the B-Box domain of
24106	PSTPIP1	17964261	MEFV and unmasking its PYD domain
24106	PSTPIP1	17964261	Enhances binding of MEFV to PYCARD
24106	PSTPIP1	14595025	Tyrosine phosphorylation of PSTPIP1 significantly enhances its interaction with MEFV
25289	CTSH	18762176	Cathepsins: key modulators of cell death and inflammatory responses An E3 ubiquitin ligase that ubiquitinates the viral 3C
27924	TRIM22	19218198	protease and acts as a component of an antiviral pathway induced by IFN against picornaviruses A member of the SWI/SNF family of proteins and is essential for the multiple changes in gene expression
28592	SMARCA4	19144648	that occur during differentiation Required for maintaining expression of several smooth
28592	SMARCA4	19342595	muscle-specific genes Part of the NLRP (Nucleotide-binding oligomerization domain, Leucine rich Repeat and Pyrin domain containing) family, has a role in apoptosis and
29985	NLRP14	18648497	inflammation and several NLRPs have been indicated as being involved in reproduction as well Heterodimerizes with CDK9 to form the positive transcriptional elongation factor b (P-TEFb) and plays a role in the activation of a subset of NF-kappaB
30254	CCNT1	18728388	dependent targets
31021	MAP2K4	10715136	Activates JNK proteins
31021	MAP2K4	11343802	Activated by MAP3K 2,3,4,5 (MEKK 2,3,4,5)
31021	MAP2K4	9841871	Activated by MAP3K 2,3,4,5 (MEKK 2,3,4,5)
31021	MAP2K4	8940179	Activated by MAP3K 2,3,4,5 (MEKK 2,3,4,5)
35553	MAP2K3	7839144	Activates MAPK14 (p38) MAP kinases
35701	SUGT1	17420470	Positively regulates NOD1 activation Bridges the HSP90 molecular chaperone system to the substrate-specific arm of SCF ubiquitin ligase complexes, suggesting a role in SCF assembly and regulation, and providing multiple complementary
35701	SUGT1	18818696	routes for ubiquitination of Hsp90 client proteins Negatively regulates adaptor protein TICAM1-
36569	SARM1	16964262	dependent Toll-like receptor signaling

			A point mutation in the murine Hem1 gene reveals an essential role for Hematopoietic Protein 1 in
38002	NCKAP1L	19015308	lymphopoiesis and innate immunity-T development is disrupted in Hem-1 deficient mice Forms a complex with ELANE (neutrophil elastase 2)
38974	CD63	18930046	and plays a role in the targeting of ELANE to primary granules in neutrophils A member of the SWI/SNF family of proteins and is
44200	SMARCA2	19144648	essential for the multiple changes in gene expression that occur during differentiation Required for maintaining expression of several smooth
44200	SMARCA2	19342595	muscle-specific genes A tyrosine protein kinase that regulates TLR4 induced
46340	BMX	18025155	IL6 in macrophages independent of MAPK14 (p38- alpha) and NF-kappaB
46853	SERPING1	17709141	A likely regulator of MASP2
			A novel cysteine-rich secreted protein associated with
49110	RETNLB	10921885	pulmonary inflammation
49787	NFKBIB	9346485	Inhibitor of NF-kappaB, binds to both NF-kappaB subunit nuclear localization signals and NFkappaB Following TNF stimulation, PRKCE phosphorylates
50152	PRKCE	19150425	TRAF2 leading to CHUK (IKK alpha) and IKBKB (IKK beta) recruitment to the TNF receptor IL-2 and IL-7 induced heterodimerization of STAT5
50532	STAT5B	9398404	isoforms in human peripheral blood T lymphoblast IL-2 and IL-7 induced heterodimerization of STAT5
50625	STAT5A	9398404	isoforms in human peripheral blood T lymphoblast
50628	SOCS5	15590694	SOCS4 and SOCS5 regulate EGFR signaling
			Essential for the development of innate and T-cell-
53133	REL	18523276	induced colitis through its ability to modulate expression of IL-12/23 family members Negatively regulates TLR-pathway driven inflammation by preventing the binding of
			phosphorylated transcription factors CREB and ATF1
53886	RPS6KA4	18690222	to IL-10 and DUSP1 promoters
85368	SH2D1A	18501771	The SLAM and SAP gene families control innate and adaptive immune responses
85368	SH2D1A	19079134	SLAM receptors and SAP influence lymphocyte interactions, development and function The SLAM and SAP gene families control innate and
103936	SLAMF8	18501771	adaptive immune responses The SLAM and SAP gene families control innate and
103966	SLAMF9	18501771	adaptive immune responses The SLAM and SAP gene families control innate and
104051	SLAMF6	18501771	adaptive immune responses SLAM receptors and SAP influence lymphocyte
104051	SLAMF6	19079134	interactions, development and function The SLAM and SAP gene families control innate and
104088	SLAMF7	18501771	adaptive immune responses SLAM receptors and SAP influence lymphocyte
104088	SLAMF7	19079134	interactions, development and function

				C2 is part of the classical and lectin complement
	301101	C2	12791093	pathways
				C2 molecule binds to C4B and is cleaved by C1S protease into C2A and C2B fragments and the resulting
				C4B2A complex (C3 convertase) is the active protease
	301101	C2	11044372	which cleaves C3
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