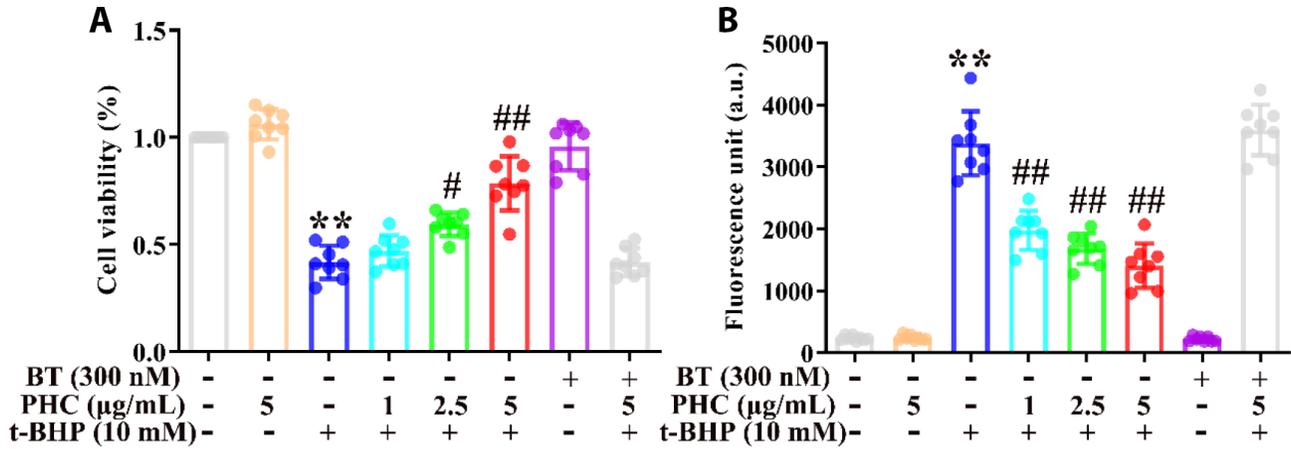
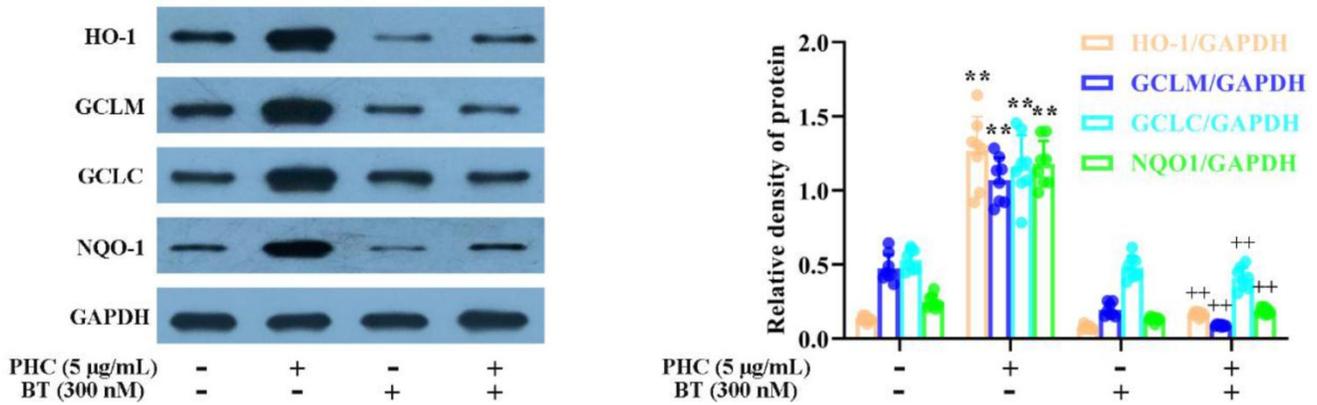


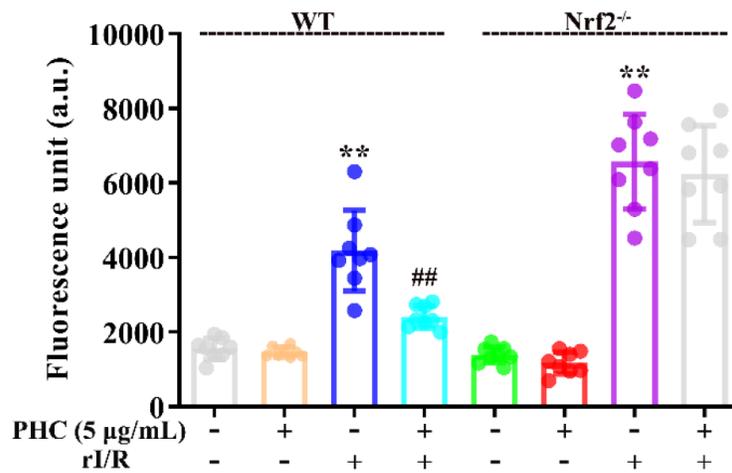
SUPPLEMENTARY FIGURES



Supplementary Figure 1. Effects of PHC and brusatol on t-BHP-induced oxidative damage in NR8383 cells. (A) NR8383 cells were stimulated with PHC (1, 2.5, 5 µg/mL) for 24 h with/without brusatol (300 nM), and then were treated with t-BHP (10 mM) for 4 h. A CCK8 assay was used to measure cell viability. (B) NR8383 cells were treated with PHC (1, 2.5, 5 µg/mL) for 24 h with/without brusatol, stained with DCFH-DA (5 µM) for 40 min and then treated with t-BHP (10 mM) for 5 min to produce ROS. A fluorescence microplate reader was used to measure ROS levels. Data are presented as the mean ± S.D. (n = 8). **P* < 0.05, ***P* < 0.01 vs. the control group. #*P* < 0.05, ##*P* < 0.01 vs. the t-BHP group.



Supplementary Figure 2. Suppression of Nrf2-induced antioxidant enzymes by brusatol. NR8383 cells were stimulated with brusatol (an antagonist of Nrf2, 300 nM) for 1 h and then exposed to PHC (5 µg/mL) for 1 h before being treated with serum from rI/R rats for 24 h. Western blotting was used to measure the protein levels of GCLM, HO-1, NQO1 and GCLC. GAPDH was used as an internal control. Data are presented as the mean ± S.D. (n = 8). **P* < 0.05, ***P* < 0.01 vs. the control group. +*P* < 0.05, ++*P* < 0.01 vs. the PHC alone group.



Supplementary Figure 3. Nrf2 dependence of the antioxidative effects of PHC in primary rat alveolar macrophages. Alveolar macrophages isolated from WT and Nrf2^{-/-} rats were pre-stimulated with PHC (5 μg/mL) for 1 h and then treated with serum from rI/R rats for 24 h. Subsequently, the cells were stained with DCFH-DA (5 μM) for 40 min, and a fluorescence microplate reader was used to determine ROS levels. Data are presented as the mean ± S.D. (n = 8). **P* < 0.05, ***P* < 0.01 vs. the control group. #*P* < 0.05, ##*P* < 0.01 vs. the rI/R group.