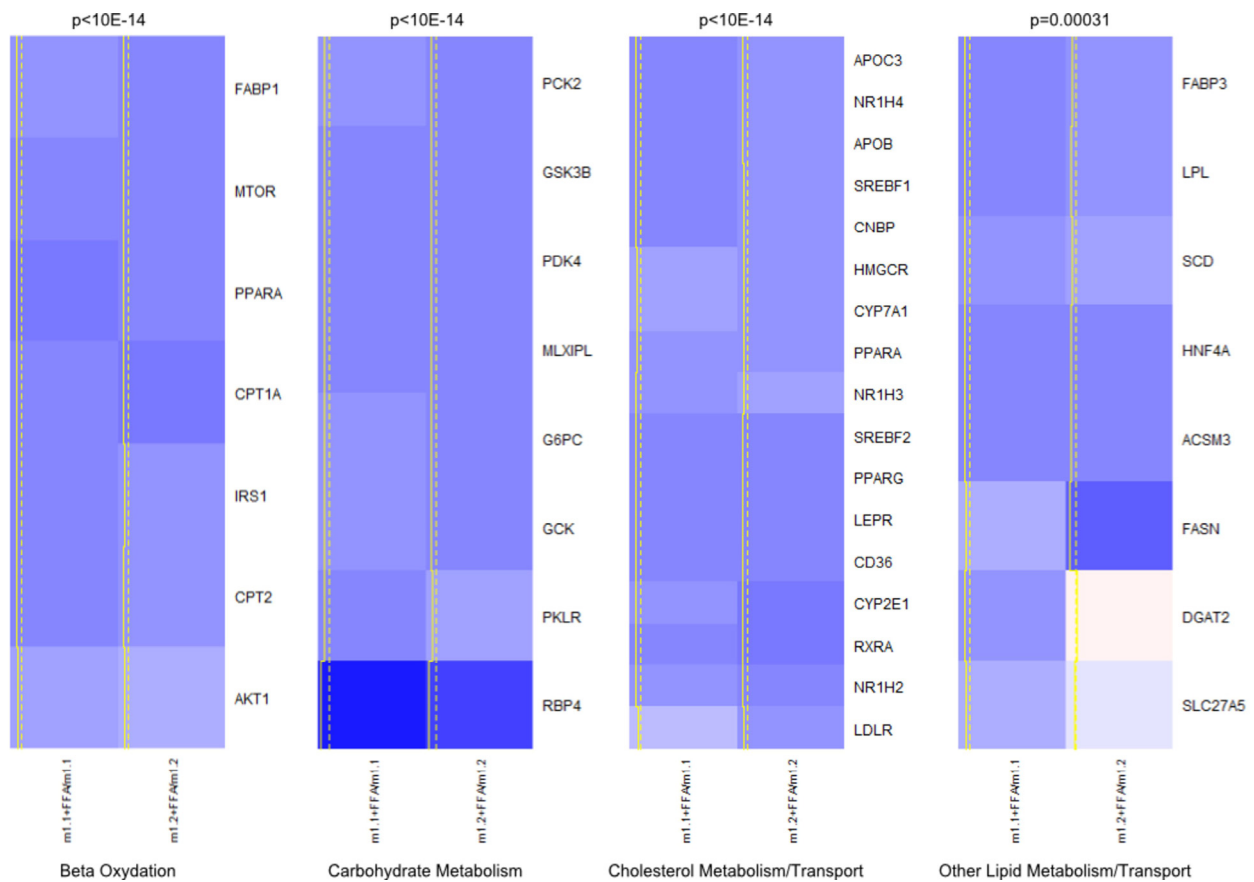


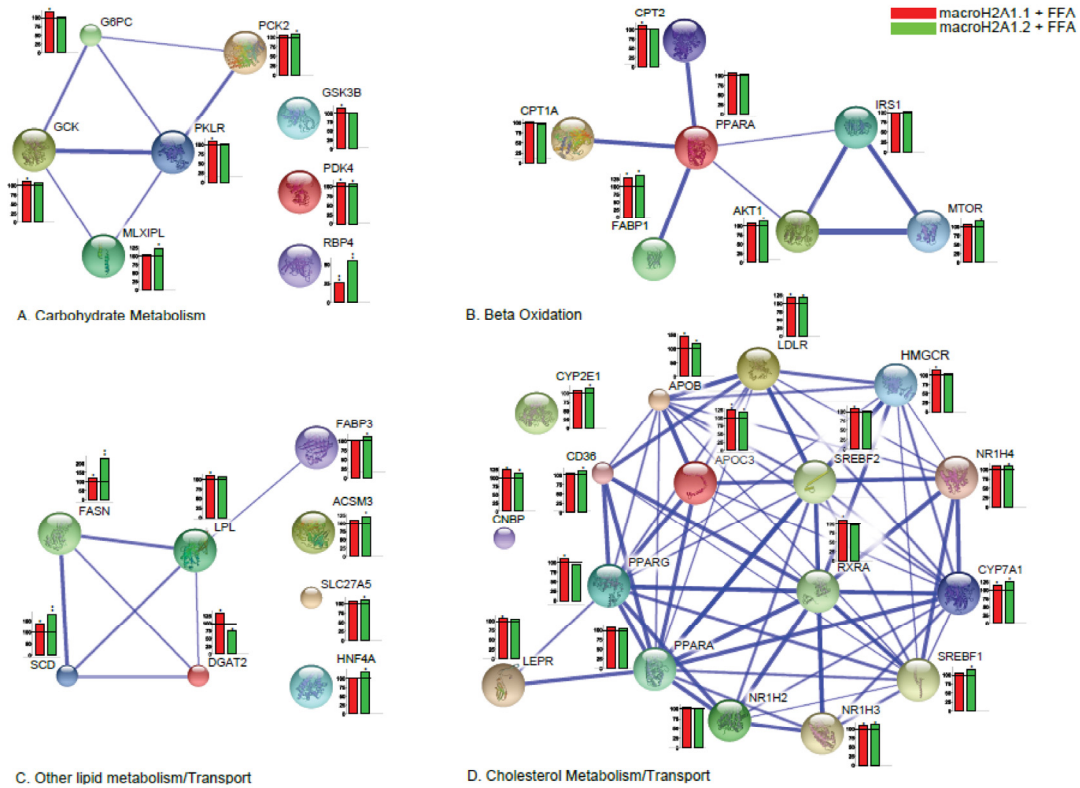
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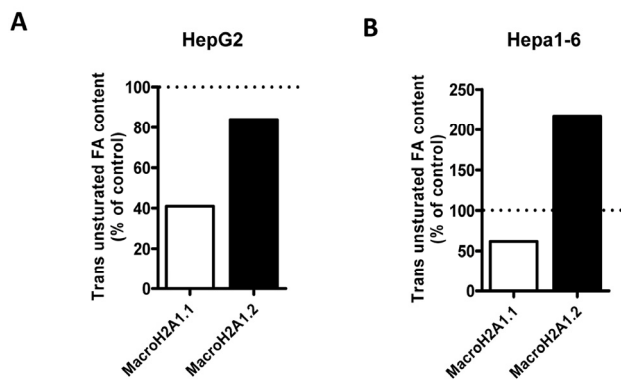
## SUPPLEMENTAL FIGURES



**Supplemental Figure 1.** Heatmap and clusters of gene expression of HepG2 cells overexpressing macroH2A1.1 (m1.1) or macroH2A1.2 (m1.2) and treated with FFA. Results are expressed as ratio of FFA-treated versus untreated cells. Optimal clusters have been computed by the pvclust method. Results were grouped in four functional processes (carbohydrate metabolism, beta-oxidation, lipid metabolism, cholesterol transport). Significance levels have been calculated via multiscale bootstrap resampling. The lower p-value of a cluster, the stronger the support of the data to the cluster. Expression levels are represented in a colour scale from blue (low expressed) to red (highly expressed) (top left).



**Supplemental Figure 2.** Differential effects of macroH2A1.1 and macroH2A1.2 on the expression of genes involved in lipid and carbohydrate metabolism in HepG2 cells. 81 genes contained in a commercially available fatty liver array were measured by qRT-PCR in HepG2 cells transiently transfected and treated with FFA, as described in the legends of Figure 1 and 2. Results were clustered in four functional hypergraphs (carbohydrate metabolism, **A**; beta-oxidation, **B**; lipid metabolism/transport, **C**; cholesterol transport/metabolism, **D**), built on a number of complementary system analyses of biological pathways, as described in the Supplemental Material & Methods section. Results of gene expression in each histogram are represented as % of FFA-treated mock-transfected (blue), FFA-treated macroH2A1.1-overexpressing (green) or FFA-treated macroH2A1.2-overexpressing (red) condition related to their respective untreated controls. Results are expressed as percentage of controls, means  $\pm$  SEM of two independent experiments. \* $p < 0.05$ .



**Supplemental Figure 3.** Overexpression of macroH2A1 isoforms (macroH2A1.1 or macroH2A1.2) and trans unsaturated fatty acids in Hepa1-6 and HepG2 cells. **A, B:** cells were transiently transfected and treated with FFA as described in the legends of Figure 1 and 2. Trans unsaturated fatty acids content was measured using TLC-chromatography in HepG2 (**A**) and Hepa1-6 (**B**) upon FFA treatment. Results are expressed as percentage of respective controls (untreated macroH2A1.1- or macroH2A1.2-overexpressing cells), and are means of two independent experiments.