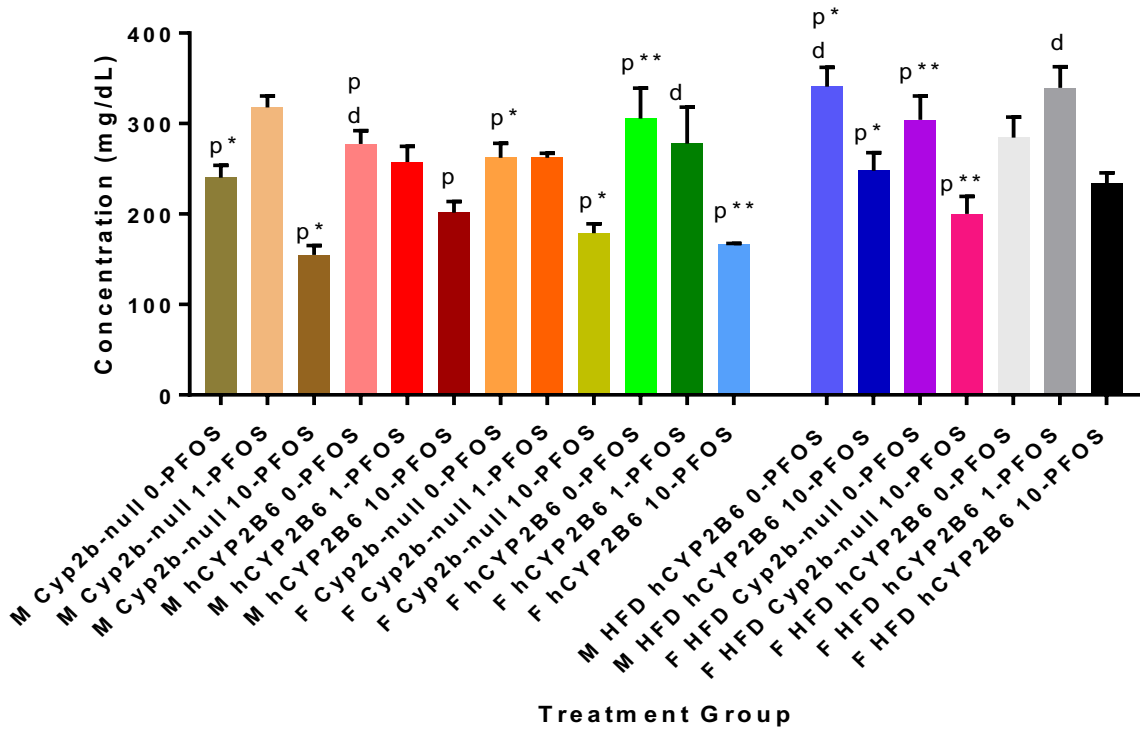


Supplemental Table 1: qPCR primer information for genes of interest

Gene	Accession Number	Forward Sequence	Reverse Sequence	Annealing Temperature (°C)
SREBP1	NM_001358315.1	5'-ACGAAGTGACACAAAAGCA-3'	5'-GCCAAAAGACAAGGGGCTAC-3'	58
CPT1A	NM_013495.2	5'-TTGATCAAGAAGTGCCGGACGAGT-3'	5'-GTCCATCATGGCCAGCACAAAGTT-3'	60
CYP2B6	KR711982.1	5'-CCATACACAGAGGCAGTCAT-3'	5'-GGTGCAGATCGATGTCTTC-3'	54.5
CD68	NM_001291058.1	5'-CGCAGACGACAATCAACCTA-3'	5'-AGTGGCATGGTGAAGAGATG-3'	59
CYP2A5	NM_007812.4	5'-CAAAGCCAAGGAAGCAAGATG-3'	5'-AGTGGTGCTGAGTGGTAATG-3'	60
CYP4A14	NM_007822.2	5'-CCTCCTCATATTGCCCTGAATAG-3'	5'-GAGTCCATAGGCCTGAGTTATT-3'	59
PPARG	NM_001127330.2	5'-TGGGTGAAACTCTGGGAGATTC-3'	5'-AATTTCTGTGAAGTGCTCATAGGC-3'	60.1
HMGCR	NM_008255.2	5'-TTAGGCATGTGGTGGTGAAG-3'	5'-GCCAAGGAGGAGCAGAATAAA-3'	60
18S	NR_003278.3	5'-ATGGCCGTTCTTAGTTGGTG-3'	5'-ATGCCAGAGTCTCGTTCGTT-3'	64
GAPDH	NM_008084.3	5'-CCTTCATTGACCTCAACTA-3'	5'-CTGGAAGATGGTGATGG-3'	50
CYP2A13	NM_000766.5	5'-TCTCCCACTTCTCCTCTGT-3'	5'-GGGAAGGAGGACAGACAATTAC-3'	59.8

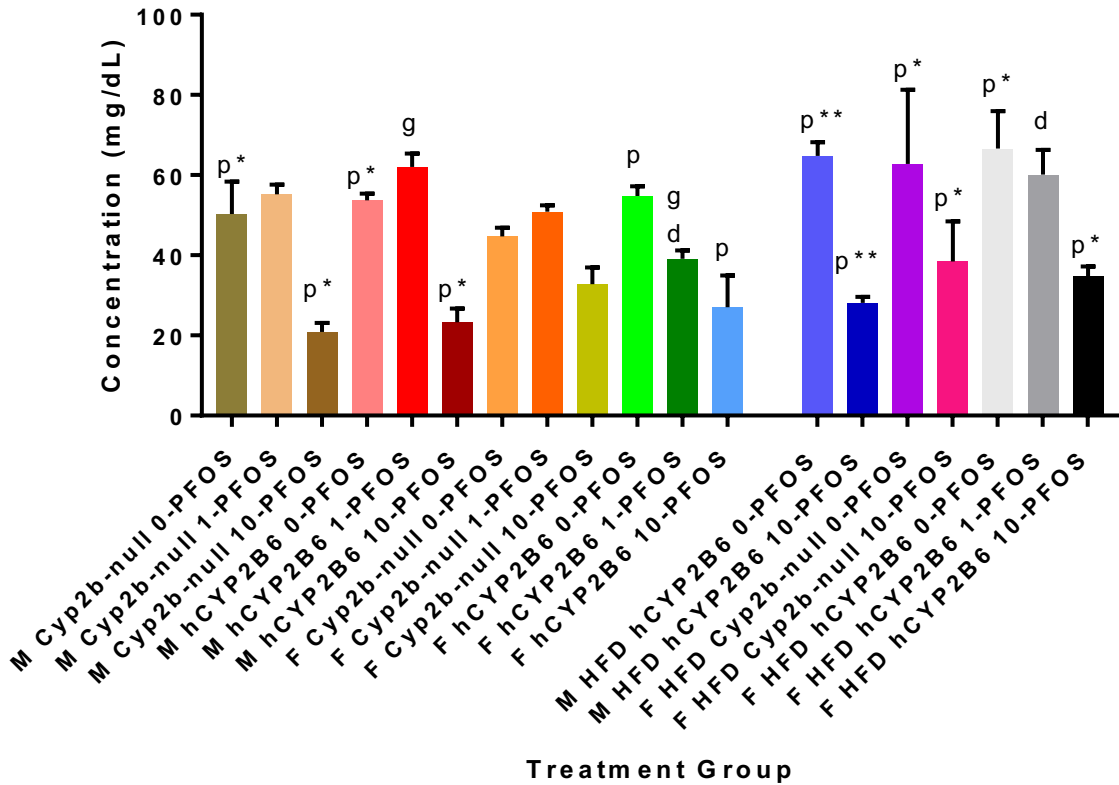
GLUCOSE



p = significant difference due to PFOS treatment
 g = significant difference due to gender
 d = significant difference due to diet
 b = significant difference due to Cyp2b

Supplemental Figure 1: Serum glucose levels (n = 5). A serum panel was performed as mentioned in the Materials and Methods and glucose levels were measured. PFOS decreased serum glucose. Data are presented as mean \pm SEM. Statistical significance was determined by one-way ANOVA followed by Fisher's LSD as the post-hoc test. A letter indicates a $p < 0.05$, letter w/ * indicates $p < 0.01$, letter w/ ** indicates $p < 0.001$ and a letter w/ *** indicates $p < 0.0001$.

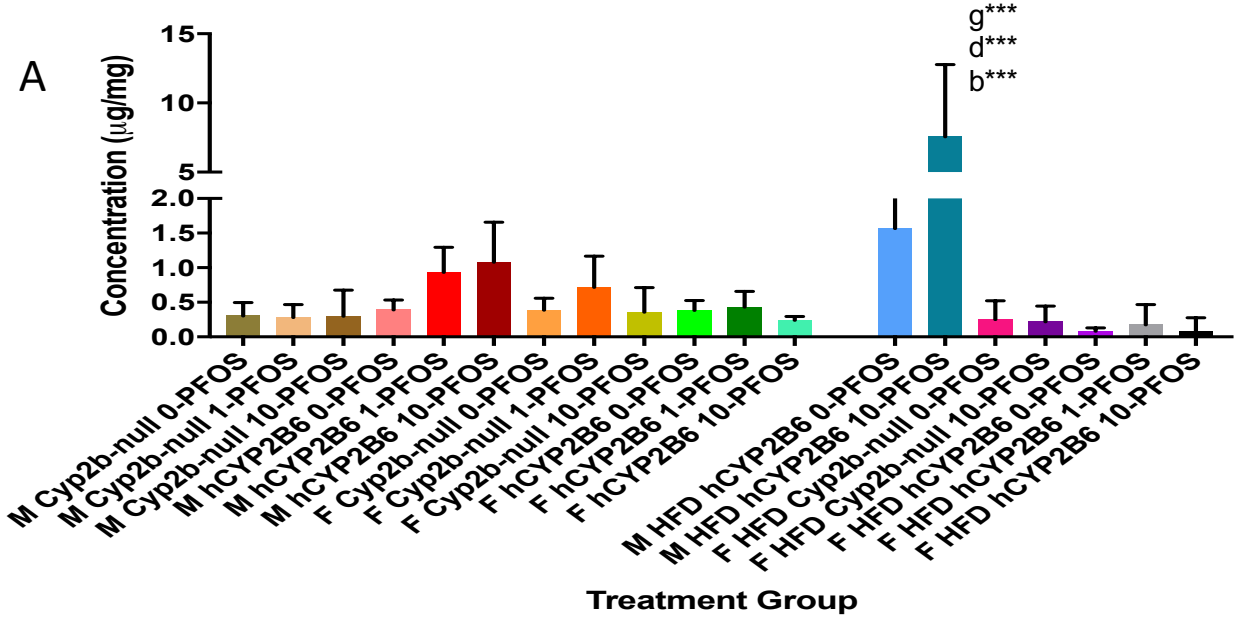
TRIG



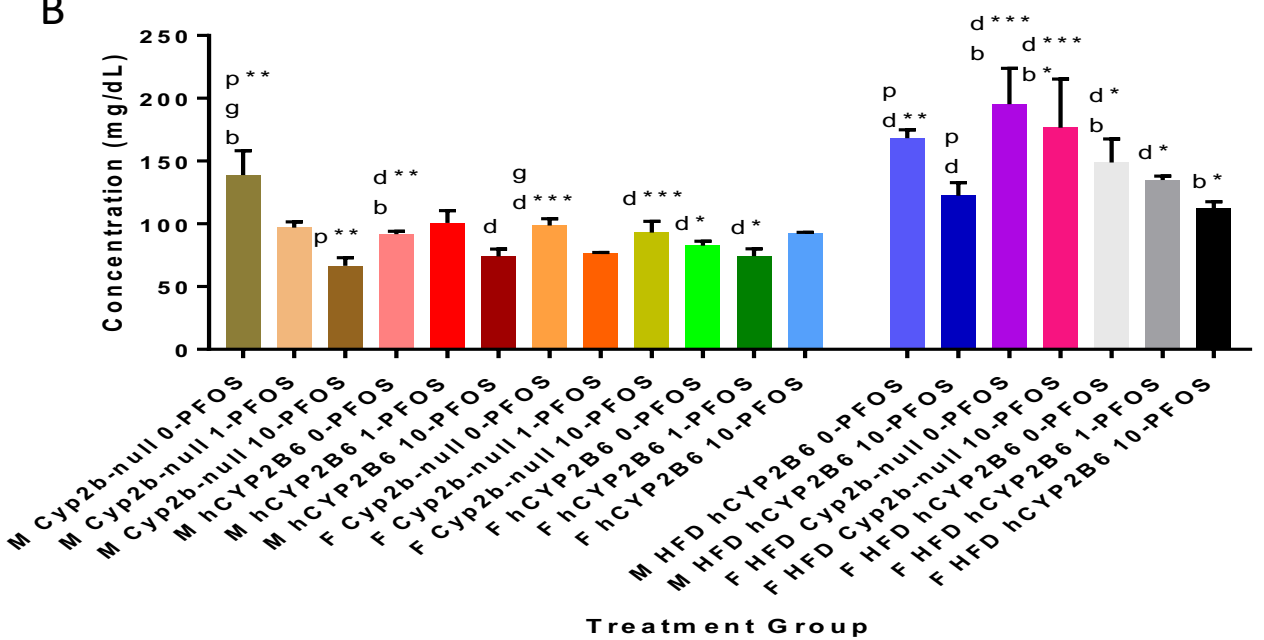
p = significant difference due to PFOS treatment
 g = significant difference due to gender
 d = significant difference due to diet
 b = significant difference due to Cyp2b

Supplemental Figure 2: Serum triglyceride levels (n=5). A serum panel was performed as mentioned in the Materials and Methods and triglyceride levels were measured. PFOS significantly decreases serum triglyceride levels. Data are presented as mean \pm SEM. Statistical significance was determined by one-way ANOVA followed by Fisher's LSD as the post-hoc test. A letter indicates a $p < 0.05$, letter w/ * indicates $p < 0.01$, letter w/ ** indicates $p < 0.001$ and a letter w/ *** indicates $p < 0.0001$.

Liver Cholesterol



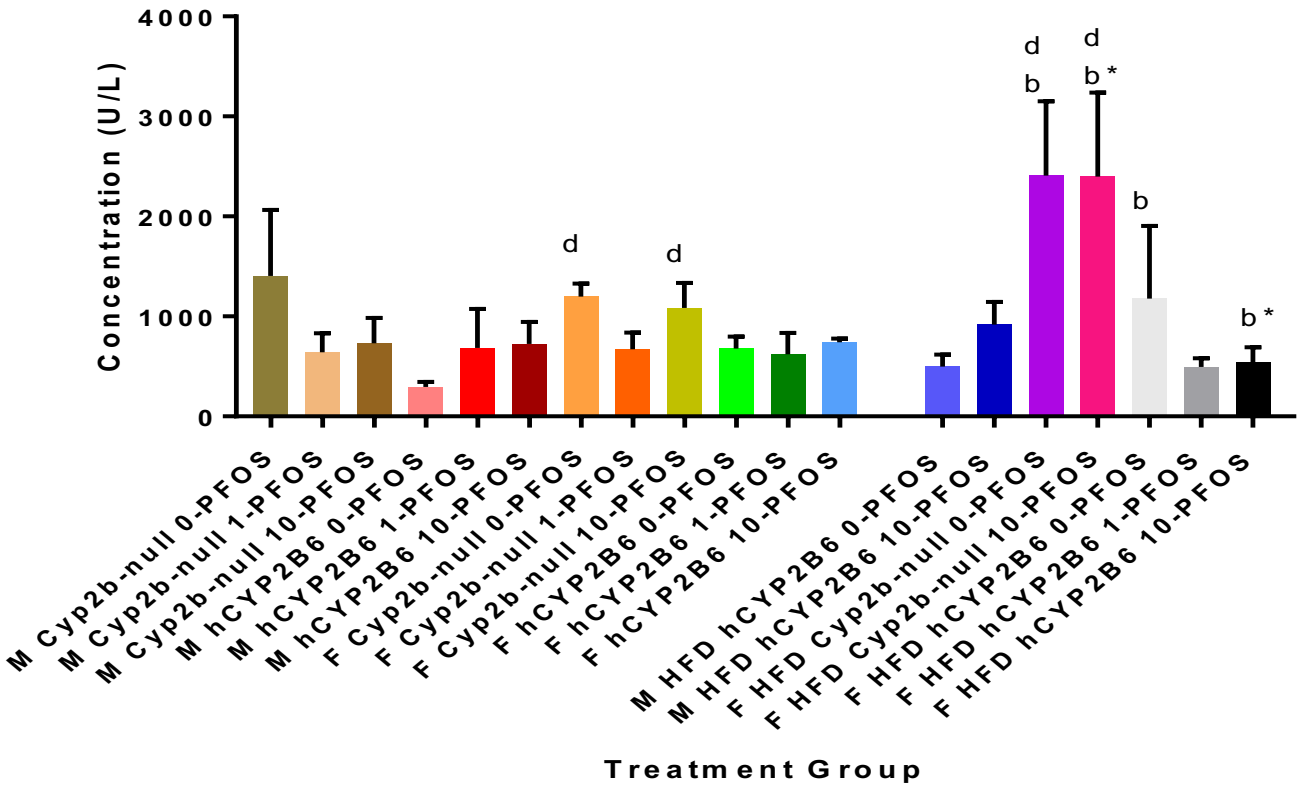
Cholesterol



p = significant difference due to PFOS treatment
g = significant difference due to gender
d = significant difference due to diet
b = significant difference due to Cyp2b

Supplemental Figure 3: Liver (A) and Serum (B) cholesterol levels (n=5). A serum panel was performed as mentioned in the Materials and Methods and cholesterol levels were measured. Increased serum cholesterol levels in the mice fed a high-fat diet with a significant increase in the cholesterol levels in Cyp2b-null mice fed a high-fat diet, regardless of PFOS concentration, compared to their hCYP2B6 counterparts. Data are presented as mean \pm SEM. Statistical significance was determined by one-way ANOVA followed by Fisher's LSD as the post-hoc test. A letter indicates a $p < 0.05$, letter w/ * indicates $p < 0.01$, letter w/ ** indicates $p < 0.001$ and a letter w/ *** indicates $p < 0.0001$.

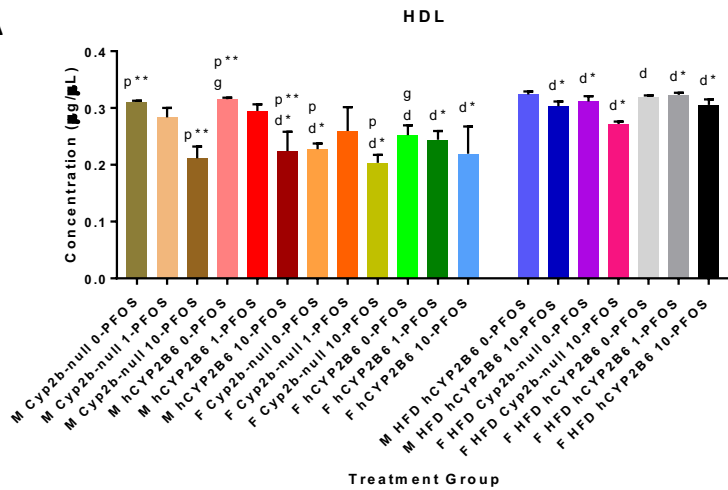
LDH



p = significant difference due to PFOS treatment
 g = significant difference due to gender
 d = significant difference due to diet
 b = significant difference due to Cyp2b

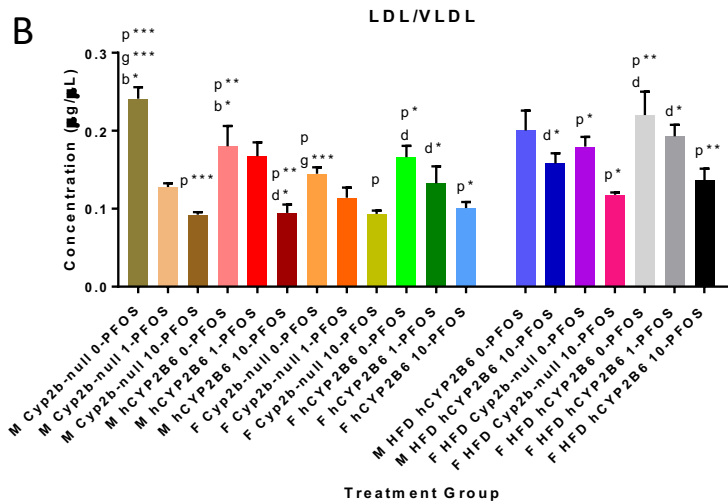
Supplemental Figure 4: Serum lactate dehydrogenase (LDH) levels (n=5). A serum panel was performed as mentioned in the Materials and Methods and LDH levels were measured. Significant increase of LDH levels in Cyp2b-null mice fed a high-fat diet regardless of PFOS concentration compared to their hCYP2B6 counterparts. Data are presented as mean \pm SEM. Statistical significance was determined by one-way ANOVA followed by Fisher's LSD as the post-hoc test. A letter indicates a $p < 0.05$, and a letter w/ * indicates $p < 0.01$.

A



p = significant difference due to PFOS treatment
 g = significant difference due to gender
 d = significant difference due to diet
 b = significant difference due to Cyp2b

B



p = significant difference due to PFOS treatment
 g = significant difference due to gender
 d = significant difference due to diet
 b = significant difference due to Cyp2b

Supplemental Figure 5: Serum HDL (A) and LDL/VLDL (B) analysis (n=4). Total serum HDL and LDL/VLDL cholesterol was determined using the HDL and LDL/VLDL Colorimetric Quantitation Assay from Sigma Aldrich (St. Louis, MO). The total HDL and LDL/VLDL cholesterol concentrations were determined according to the manufacturer's instructions. PFOS suppresses both HDL and LDL/VLDL in all mice. Data are presented as mean ± SEM. Statistical significance was determined by one-way ANOVA followed by Fisher's LSD as the post-hoc test (n=5). A letter indicates a $p < 0.05$, letter w/ * indicates $p < 0.01$, letter w/ ** indicates $p < 0.001$ and a letter w/ *** indicates $p < 0.0001$.