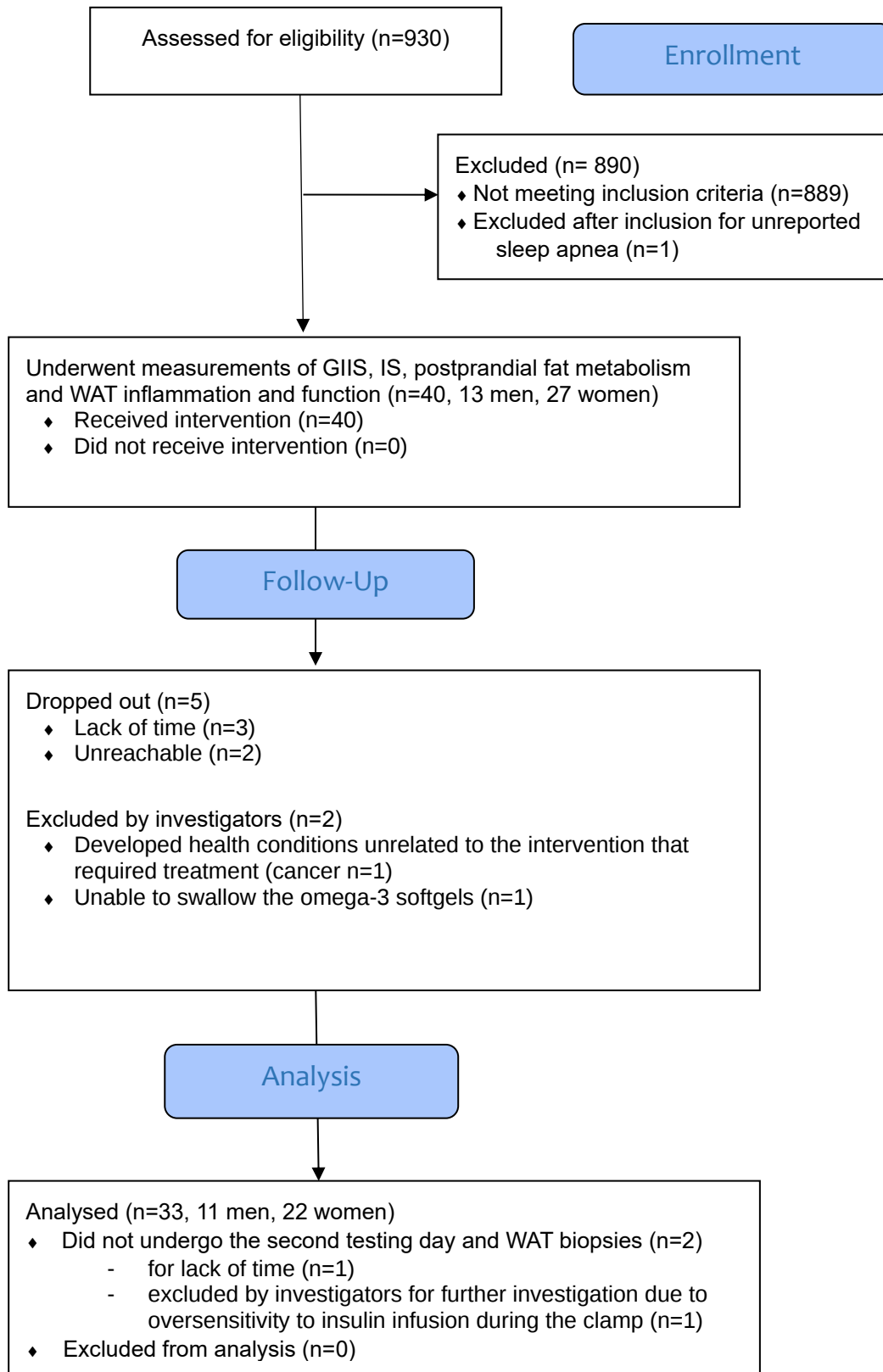


EPA and DHA inhibit LDL-induced upregulation of human adipose tissue NLRP3 inflammasome/IL-1 β pathway and its association with diabetes risk factors

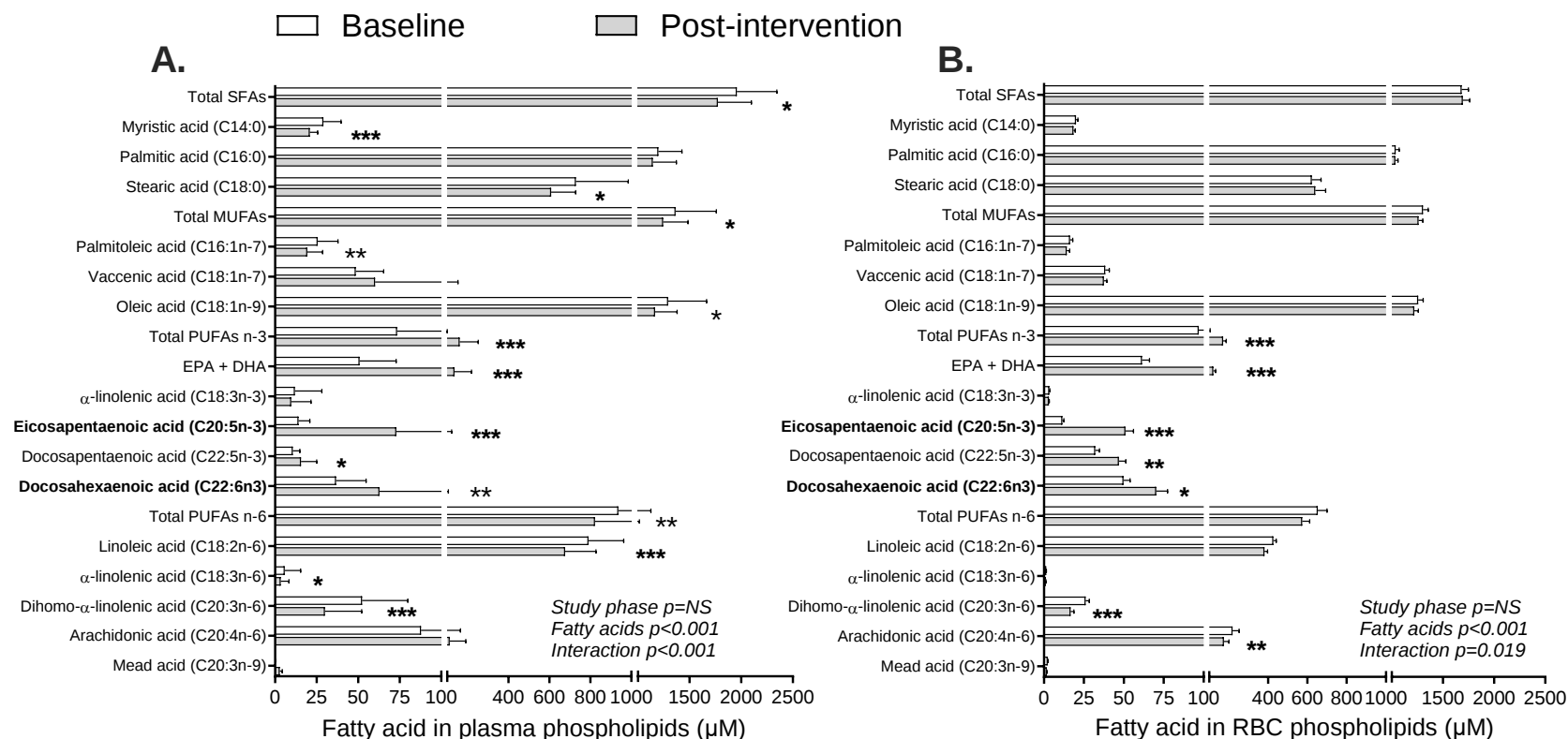
Valérie Lamantia^{1,2,3}, Simon Bissonnette^{1,2,3}, Myriam Beaudry^{1,2,3}, Yannick Cyr^{1,2,3}, Christine Des Rosiers^{1,4}, Alexis Baass^{1,2,5}, May Faraj^{1,2,3,5}

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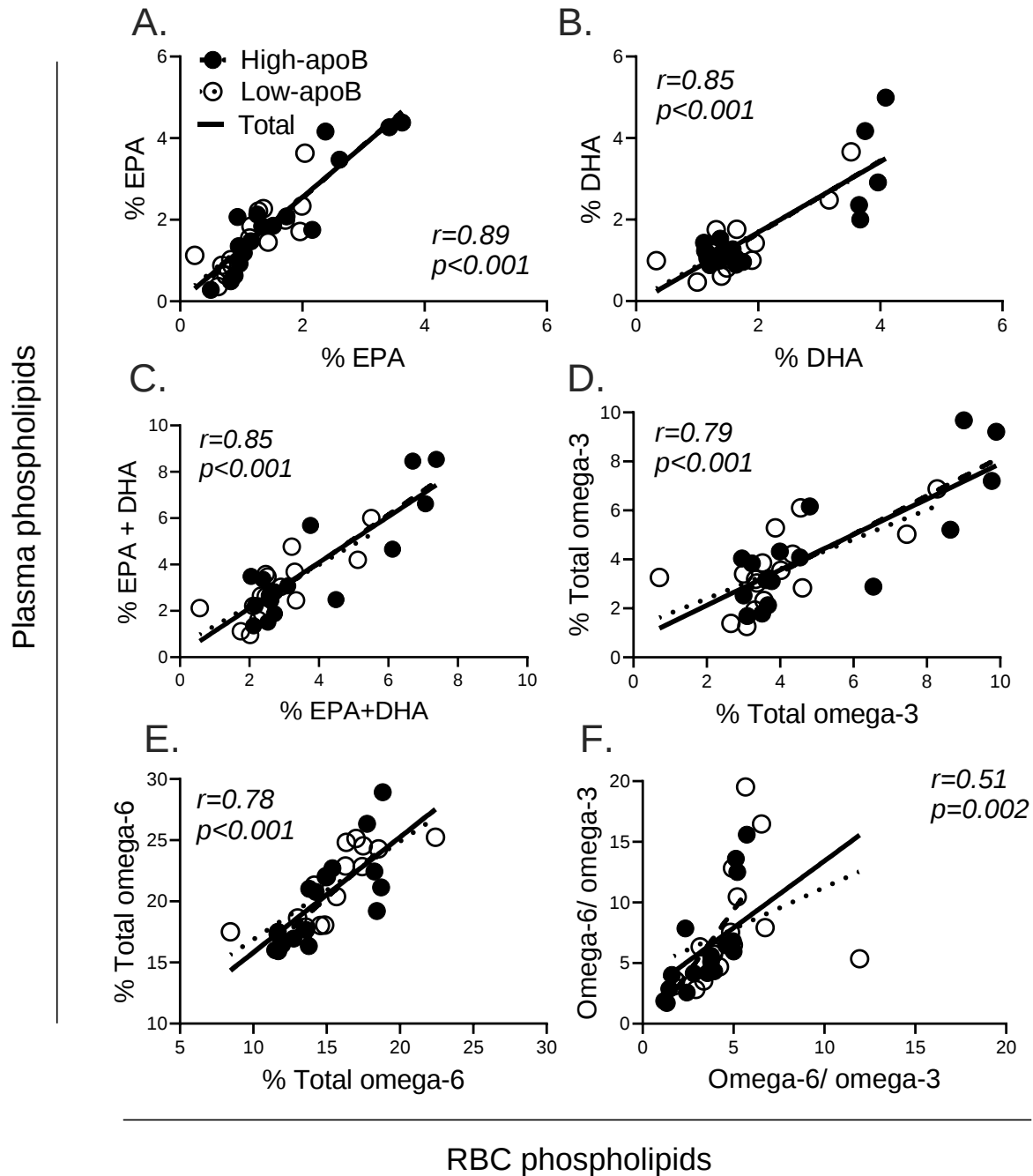
Supplemental Figure 1: Subject flow chart



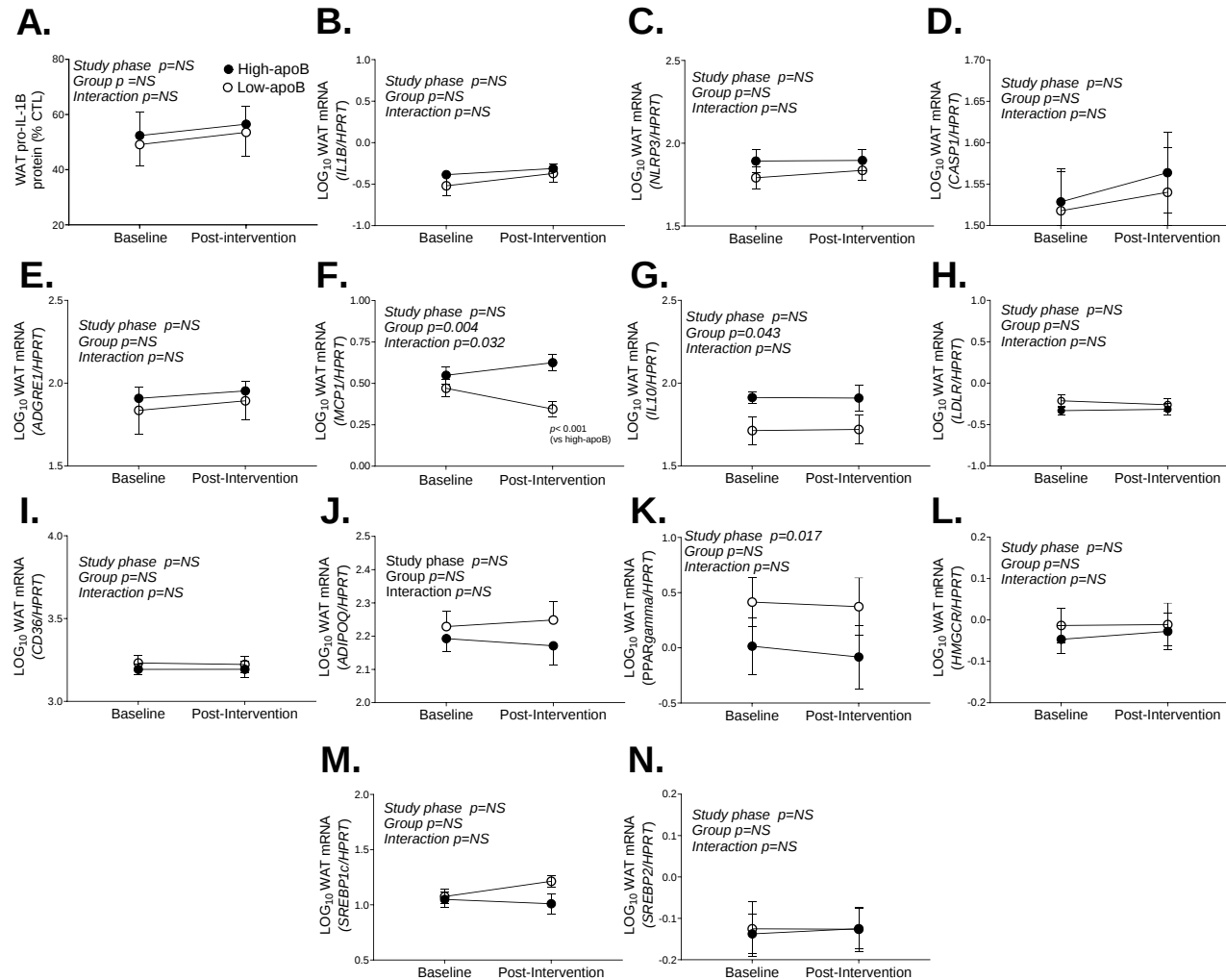
Supplemental Figure 2: Fatty acid concentrations in the phospholipid layer of A) plasma (i.e. lipoproteins) and B) red blood cells (RBC) at baseline and following the 12-week supplementation with EPA and DHA (2.7 g/d) in the 33 subjects who completed the trial. Data was analyzed by mixed-model analysis with interaction. * for $p<0.05$, ** for $p<0.01$ and *** for $p<0.001$ versus baseline. N.B. Mead acid was excluded from analysis as only N=2 subjects had measurable post-intervention concentration in RBC-PL.



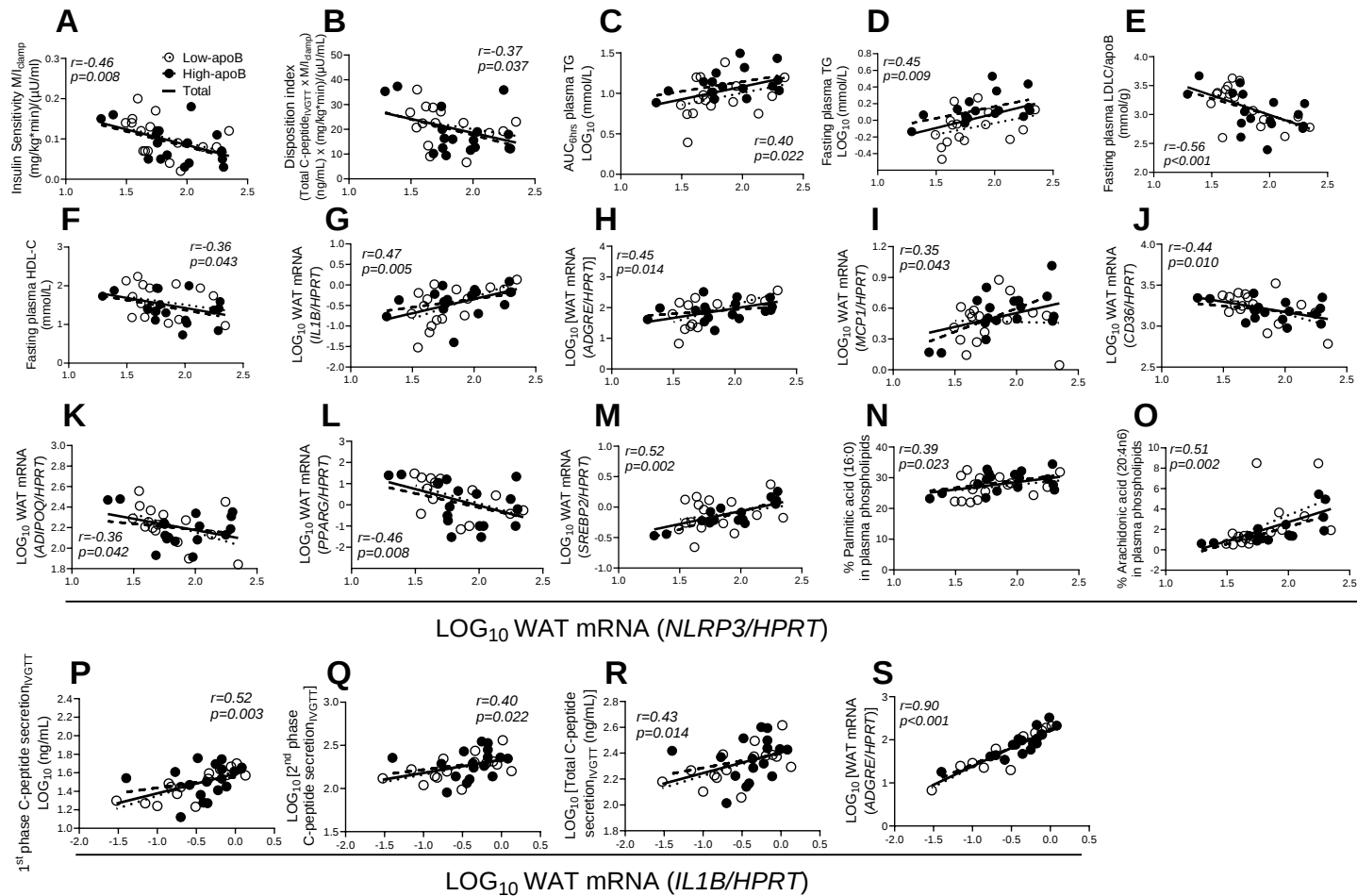
Supplemental Figure 3: Correlation between RBC and plasma phospholipids content of % EPA (A), % DHA (B), % EPA and DHA (C), % total omega-3 (D), % total omega-6 (E), and omega-6/omega-3 ratio (F) in subjects with low-apoB (N=16, open circle) and high-apoB (N=17, closed circle) who completed 12-week supplementation with EPA and DHA (2.7 g/d). Solid regression line represents correlation in all subjects. Data was analyzed by Pearson correlation as in methods.



Supplemental Figure 4: Baseline and post-intervention of WAT protein expression of pro-IL-1 β (A) and WAT mRNA expression of *IL1B* (B), *NLRP3* (C), *CASP1* (D), *ADGRE1* (E), *MCP1* (F), *IL10* (G), *LDLR* (H), *CD36* (I), *ADIPOQ* (J), *PPARG* (K), *HMGCR* (L), *SREBP1c* (M) and *SREBP2* (N) normalized to *HPRT* in subjects with low-apoB (baseline N=16, post-intervention N=14) and high-apoB (baseline N=17, post-intervention N=16) for missing data. Data analyzed by mixed-model analysis with group x time interaction.



Supplemental Figure 5: Pearson correlation of fasting baseline WAT mRNA expression of *NLRP3* normalized for *HPRT* with insulin sensitivity as M/I_{clamp} (A), total disposition index (total C-peptide $_{\text{IVGTT}} \times M/I_{\text{clamp}}$) (B), $AUC_{6\text{hrs}}$ plasma TG (C), fasting plasma TG (D), estimated fasting LDL size (E), fasting plasma HDL-C (F), and WAT mRNA expression of *IL1B* (G), *ADGRE1* (H), *MCPI* (I), *CD36* (J), *ADIPOQ* (K), *PPARG* (L), and *SREBP2* (M) normalized to *HPRT*, and % fasting plasma phospholipid palmitate (N), and arachidonate (O), and correlation of fasting baseline WAT mRNA expression of *IL1B* with 1st phase (P), 2nd phase (Q), total C-peptide secretion $_{\text{IVGTT}}$ (R), and WAT mRNA expression of *ADGRE1* normalized to *HPRT* (S) in subjects with low-apoB (N=16, open circles, dotted regression line) and high-apoB (N=17, closed circles, dashed regression line) who completed the trial. Solid regression line represents pooled data for all subjects.



Supplemental Figure 6: Pearson correlation of fasting post-intervention WAT mRNA expression of *NLRP3* normalized for *HPRT* with insulin sensitivity as M/I_{clamp} (A), total disposition index (total C-peptide_{IVGTT} x M/I_{clamp}) (B), AUC_{6hrs} plasma TG (C), fasting plasma TG (D), estimated fasting LDL size (E), fasting plasma HDL-C (F), and WAT mRNA expression of *IL1B* (G), *ADGRE1* (H), *MCPI* (I), *CD36* (J), *ADIPOQ* (K), *PPARG* (L), and *SREBP2* (M) normalized to *HPRT*, and % fasting plasma phospholipid palmitate (N), and arachidonate (O), and correlation of fasting baseline WAT mRNA expression of *IL1B* with 1st phase (P), 2nd phase (Q), total C-peptide secretion_{IVGTT} (R), and WAT mRNA expression of *ADGRE1* normalized to *HPRT* (S) in subjects with low-apoB (N=15, open circles, dotted regression line) and high-apoB (N=1, closed circles, dashed regression line) who completed the trial and have post-intervention WAT data. Solid regression line represents pooled data for all subjects.

