

Online supplemental materials

Supplemental Figure Legends

Supplemental Figure 1: ECSHIP2^{Δ/+} mice do not exhibit a pro-inflammatory state. **A)** Truncated SHIP2 mRNA was detectable in CD11b⁺ myeloid cells from ECSHIP2^{Δ/+}, although >3000-fold lower level than in lung EC (n=4,4,2). **B)** Blood leukocyte populations were comparable in ECSHIP2^{Δ/+} and control littermates (n≥8). **C)** Peripheral blood mononuclear cell SHIP2 activity was not altered in ECSHIP2^{Δ/+} (n=6,7). **D,E)** Serum TNFα and IL-6 were comparable in ECSHIP2^{Δ/+} and control littermates (n=8,14). **F)** ECSHIP2^{Δ/+} had similar TNFα, IL-6, and IL-1β mRNA expression in white adipose tissue and skeletal muscle (n≥4). **G)** White adipose tissue collagen staining with Sirius Red, a feature of chronic inflammation, was similar in ECSHIP2^{Δ/+} and control littermates (n=6,11).

Supplemental Figure 2: Juvenile ECSHIP2^{Δ/+} mice have normal EC expression of key signaling nodes and vascularization of metabolic tissues. **A)** The increased expression of Akt, Rictor and eNOS seen in 10-month old ECSHIP2^{Δ/+} mice was not apparent in 6-week old mice (n≥5). **B,C)** 6-week old ECSHIP2^{Δ/+} mice exhibited normal capillary density in white adipose tissue and skeletal muscle (n=5,4).

Supplemental Figure 3: Insulin receptor and insulin receptor substrate 1/2 signaling is unchanged in ECSHIP2^{Δ/+} mice. **A)** Basal expression and phosphorylation of the insulin receptor and insulin receptor substrate 1/2 were comparable in ECSHIP2^{Δ/+} mice and littermate controls (n≥5). **B)** Insulin-stimulated phosphorylation of the insulin receptor and insulin receptor substrate 1/2 were comparable in ECSHIP2^{Δ/+} mice and littermate controls (n≥5).