SUPPLEMENTARY DATA

Supplementary 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).



SUPPLEMENTARY DATA

Supplementary Figure 2. Juvenile ECSHIP2^{$\Delta/+$} 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^{$\Delta/+$} mice was not apparent in 6-week old mice (n \geq 5). B,C) 6-week old ECSHIP2^{$\Delta/+$} mice exhibited normal capillary density in white adipose tissue and skeletal muscle (n=5,4).



SUPPLEMENTARY DATA

Supplementary Figure 3. Insulin receptor and insulin receptor substrate 1/2 signaling is unchanged in ECSHIP2^{$\Delta/+$} mice. A) Basal expression and phosphorylation of the insulin receptor and insulin receptor substrate 1/2 were comparable in ECSHIP2^{$\Delta/+$} mice and littermate controls (n \geq 5). B) Insulin-stimulated phosphorylation of the insulin receptor and insulin receptor substrate 1/2 were comparable in ECSHIP2^{$\Delta/+$} mice and littermate controls (n \geq 5). B) Insulin-stimulated phosphorylation of the insulin receptor and insulin receptor substrate 1/2 were comparable in ECSHIP2^{$\Delta/+$} mice and littermate controls (n \geq 5).

