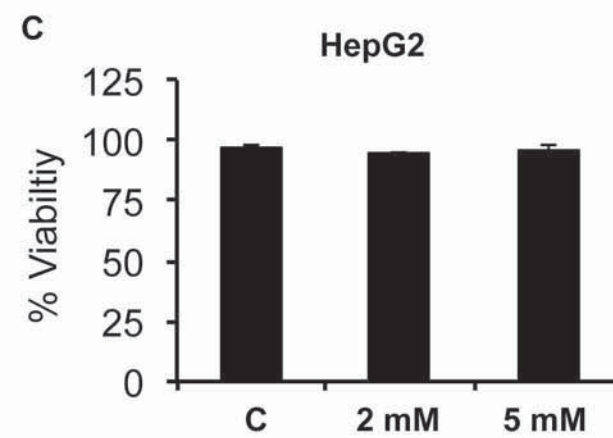
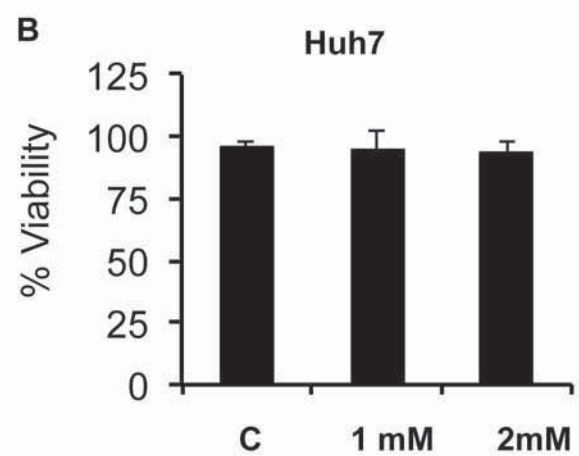
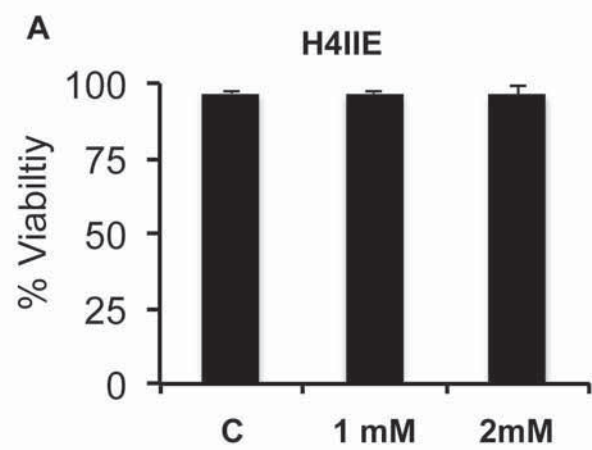
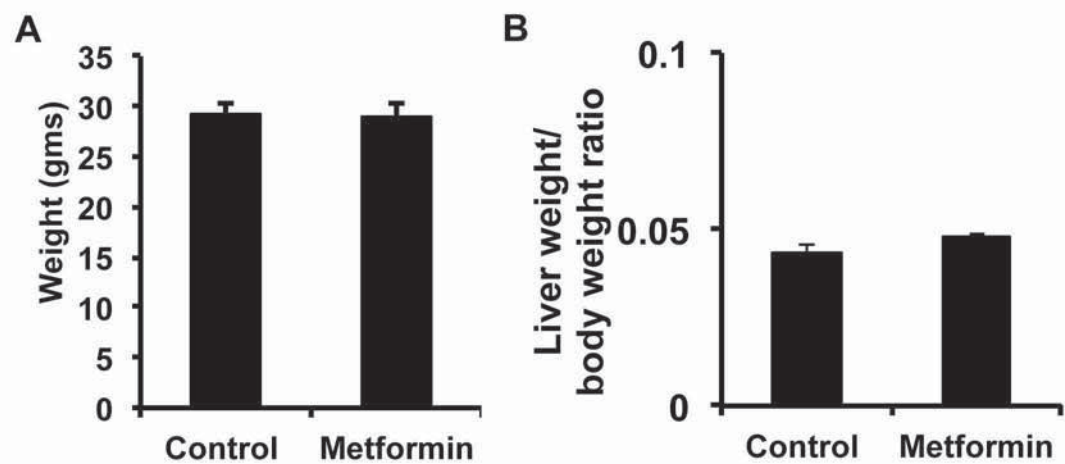


Supplemental figure 1

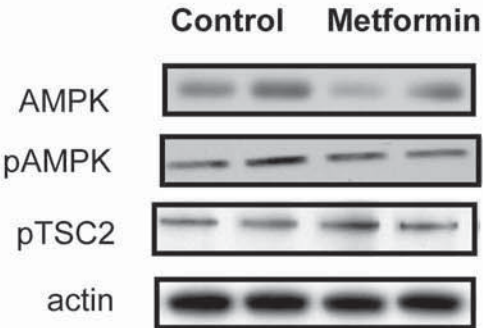


Supplemental figure 2

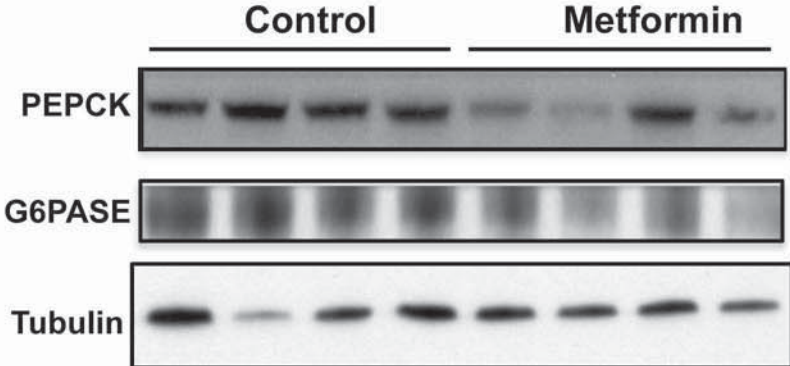


Supplemental figure 3

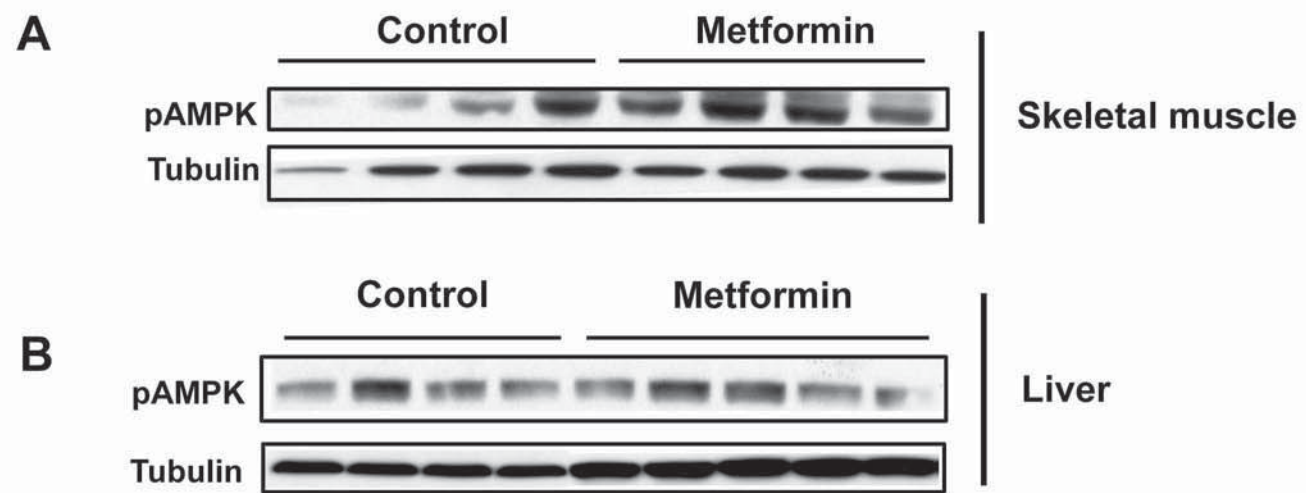
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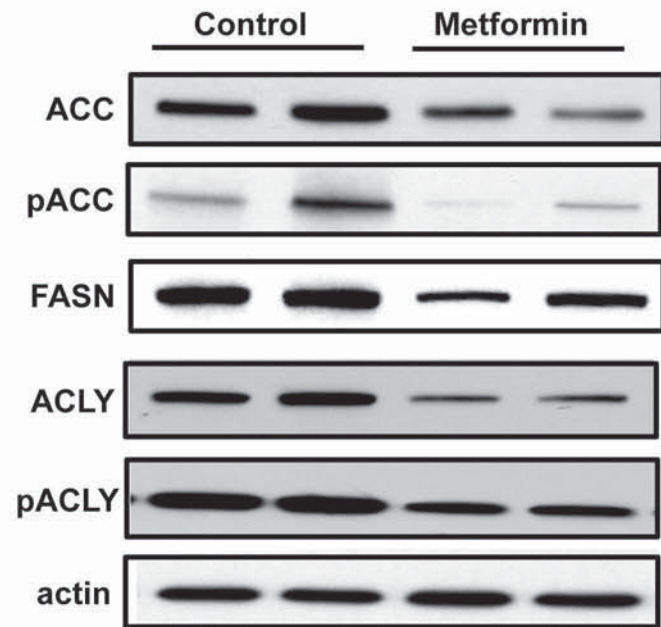
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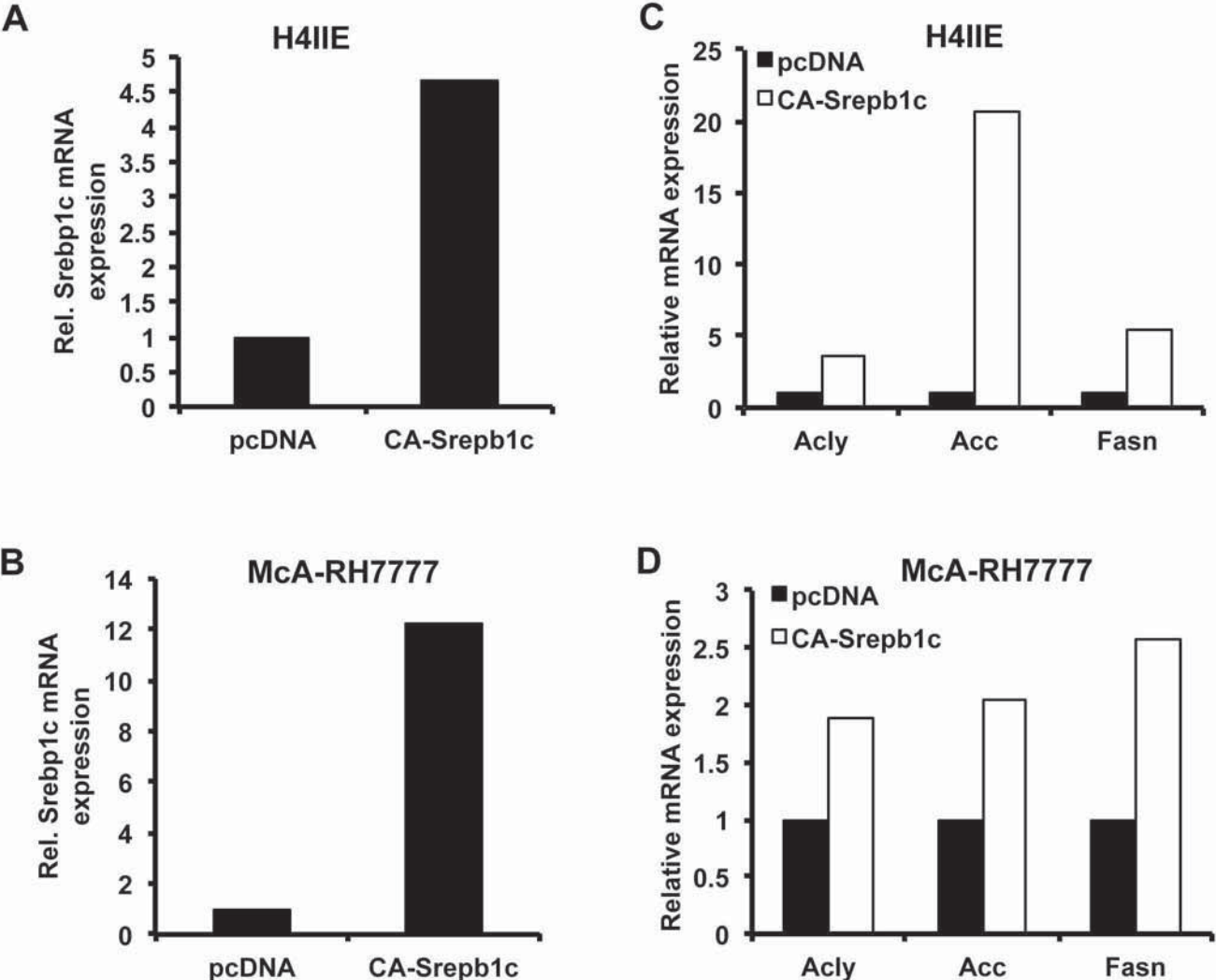
Supplemental figure 4



Supplemental figure 5



Supplemental figure 6



### **Supplemental figure legends**

**Supplemental Figure 1.** Metformin does not reduce cell viability. A) H4IIE, B) MCA-RH7777, C) Huh7 and D) HepG2 cells were treated with indicated doses of metformin for 48 hrs. Cell viability was determined using the Countess Automated Cell Counter with trypan blue as described by manufacturer (Invitrogen). N=3 ± SD.

### **Supplemental Figure 2.**

Metformin does not alter A) body weight or B) liver weight/body weight ratio of treated mice. Control fed and metformin fed mice (250 mg/kg) were weighed after 24 weeks following DEN treatment. Livers were removed and weighed and compared to total weight (g). N=4-6 ± SD.

### **Supplemental Figure 3.**

Metformin does not alter AMPK activation in the liver but reduces gluconeogenic enzyme expression. A) Additional samples were examined for AMPK, pAMPK and phosphorylated TSC2. B) Liver lysates from control and metformin treated mice were immunoblotted for PEPCK, G6Pase and tubulin as a loading control.

### **Supplemental figure 4.**

Metformin induces AMPK activation in muscle but not liver. Mice were treated with metformin for two weeks and liver and muscle tissue removed. Protein lysates from A) skeletal muscle and B) liver were immunoblotted for phosphorylated AMPK. Tubulin was used as a loading control.

**Supplemental figure 5.**

Metformin reduces lipogenic enzyme expression and activity. Additional samples of protein lysates from liver of control and metformin treated mice were immunoblotted as indicated and as described in materials and methods.

**Supplemental figure 6.**

Ectopic expression of Srebp1c promotes Acly, ACC and Fasn expression. H4IIE and McA-RH7777 cells stably expressing vector control pcDNA or CA-Srebp1c were established as described in materials and methods. RNA was isolated and RTPCR performed for Srebp1c, ACLY, ACC and FASN.