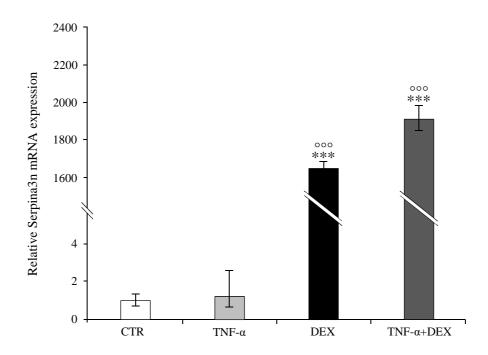
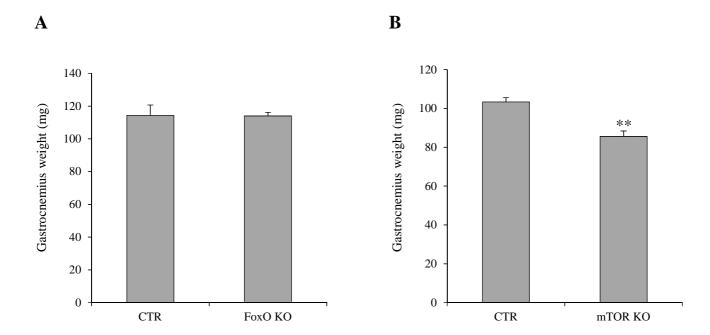


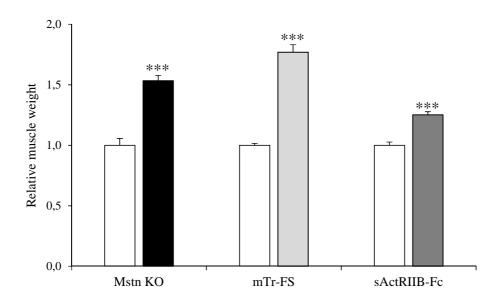
**Figure S1** Cachexia, LPS-induced sepsis and STZ-induced diabetes are associated with muscle atrophy. Gastrocnemius weight of cachectic vs CTR mice (A, n=8/group), LPS vs CTR mice (B, n=6/group), and STZ or STZ+INS vs CTR rats (C, n=6/group). Results are means  $\pm$  SE. Statistical analysis was performed using unpaired t-test or 1-way ANOVA and Tukey posttest (\*\*\* p<0.001 vs CTR,  $^{\circ\circ\circ}$  p<0.001 vs STZ+INS).



**Figure S2** TNF-α does not induce neither significantly potentiate the GC-induced expression of Serpina3n. Serpina3n mRNA levels of  $C_2C_{12}$  cells after 48h of dexamethasone (10<sup>-6</sup>M) and/or TNF-α (10ng/ml) treatment. Results are means  $\pm$  SE. Statistical analysis was performed using 1-way ANOVA and Tukey posttest (\*\*\* p<0.001 vs CTR, °°° p<0.05 vs TNF-α).



**Figure S3** mTOR inhibition, but not FoxO, are associated with muscle atrophy. Gastrocnemius weight of FoxO KO vs CTR mice (A, n=3/group), and mTOR KO vs CTR mice (B, n=3 and 5/group respectively). Results are means  $\pm$  SE. Statistical analysis was performed using unpaired t-test (\*\* p<0.01 vs CTR).



**Figure S4** Muscle hypertrophy caused by Myostatin (Mstn) inhibition. Weight of GC muscle in Mstn KO and WT mice (n=6/group), and of TA in mTr-FS (Follistatin) and WT mice (n=4/group) and in sActRIIB-treated and saline-treated mice (n=6/group). Results are means  $\pm$  SE. Statistical analysis was performed using unpaired t-test (\*\*\* p<0.001 vs CTR or WT).

## **Supplementary method**

## TNF- $\alpha$ treatment of $C_2C_{12}$ myotubes

To assess the effects of TNF- $\alpha$  treatment on Serpina3n expression, differentiated  $C_2C_{12}$  cells were treated with recombinant mouse TNF- $\alpha$  (10ng/ml, R&D Systems, Abingdon, UK) with or without DEX at  $10^{-6}$  M for 48h in 2% HS differentiation medium.