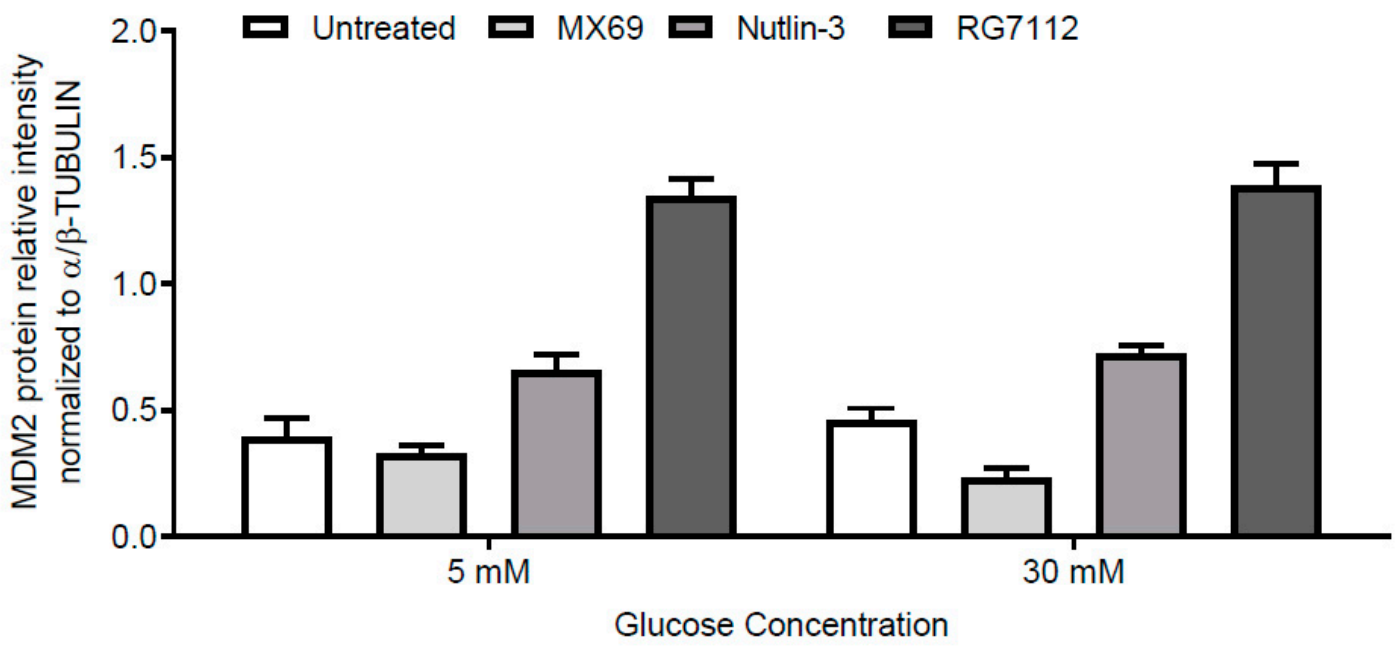
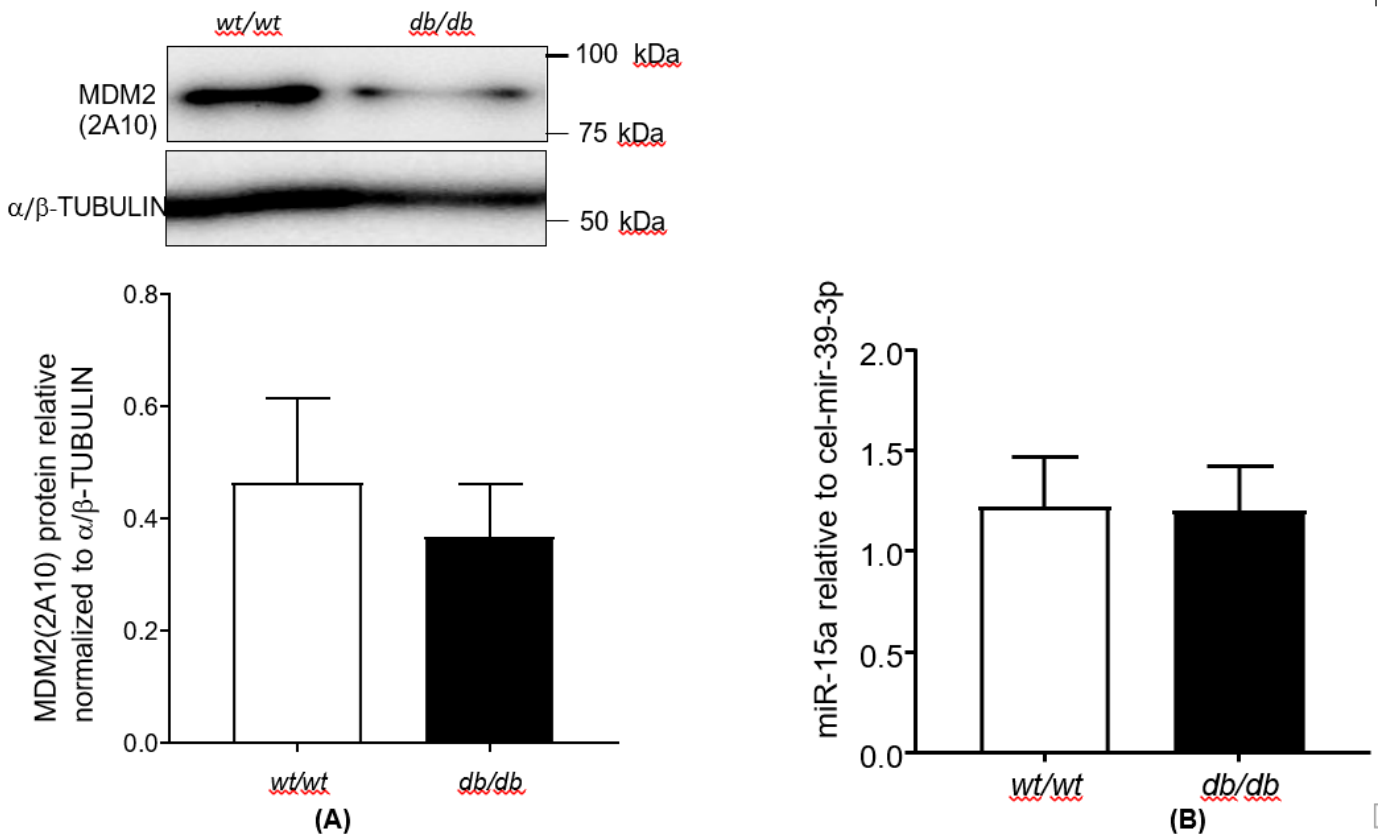


Supplementary Figure S1. Human dermal microvascular endothelial cells (HDMECs) were treated with 30 mM or 5 mM glucose and Murine Double Minute-2 (MDM2) inhibitors (MX69, Nutlin-3 and RG7112) for 6 h. Levels of MDM2 protein were measured by western blot. α,β-TUBULIN was used as a loading control. Quantification shows means ± SEM, $n = 6$. Two-way ANOVA shows an overall effect of MDM2 activity inhibition, $p < 0.0001$. Nutlin-3 and RG7112 treated cells presented significant higher level of MDM2 than untreated cells. MX69 treated cells showed a lower level of MDM2 than un-treated cells.



Supplementary Figure S2. Level of MDM2 protein and miR-15a are similar in the gastrocnemius skeletal muscle of wild type and *db*/*db* mice aged 13 weeks. (**A**) MDM2 protein levels in *wt*/*wt* and *db*/*db* mice as measured by 2A10 antibody. α,β-TUBULIN was used as a loading control. (**B**) Mature miR-15a level relative to exogenous spike-in cel-miR-39-3p. Data are means ± SEM (*wt*/*wt* mice $n = 8$ and *db*/*db* mice $n = 6$).