

Effects of Overexpression of Neurosecretory Protein GL-Precursor Gene on Glucose Homeostasis and Insulin Sensitivity in Mice

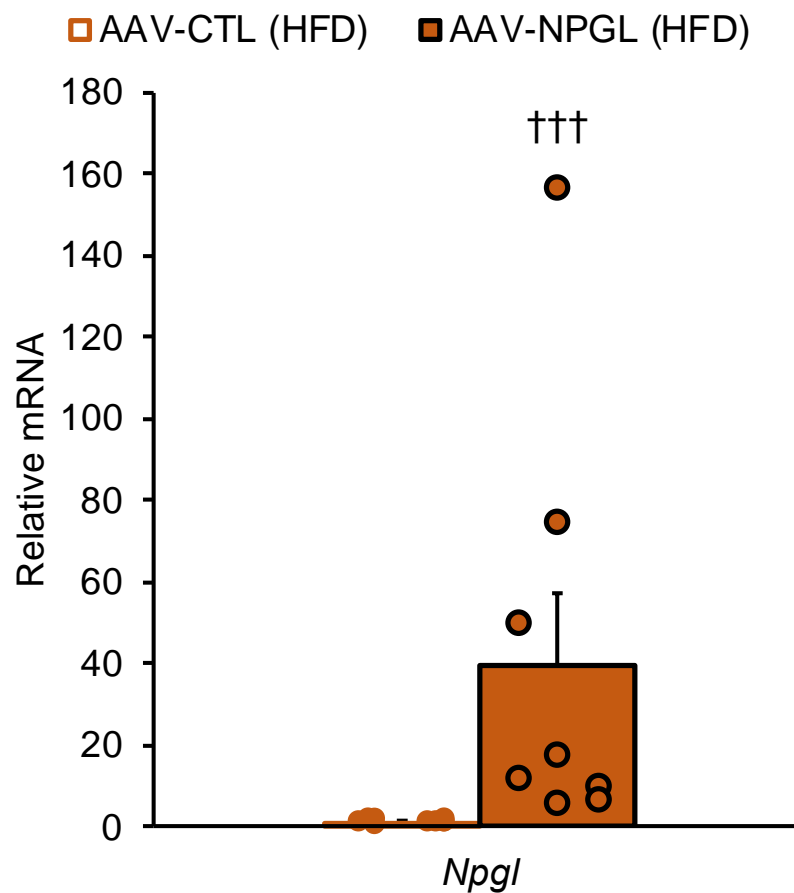


Figure S1. Efficacy of *Npgl* overexpression in the mediobasal hypothalamus at the experimental endpoint. The panel shows mRNA expression level of *Npgl* obtained upon injection the AAV-based control vector (AAV-CTL) or the AAV-based NPGL-precursor gene vector (AAV-NPGL) in high-fat diet (HFD)-fed mice. Circles represent individual data points. Each value represents the mean \pm standard error of the mean ($n = 8$; ††† $p < 0.005$ for Mann-Whitney U test).

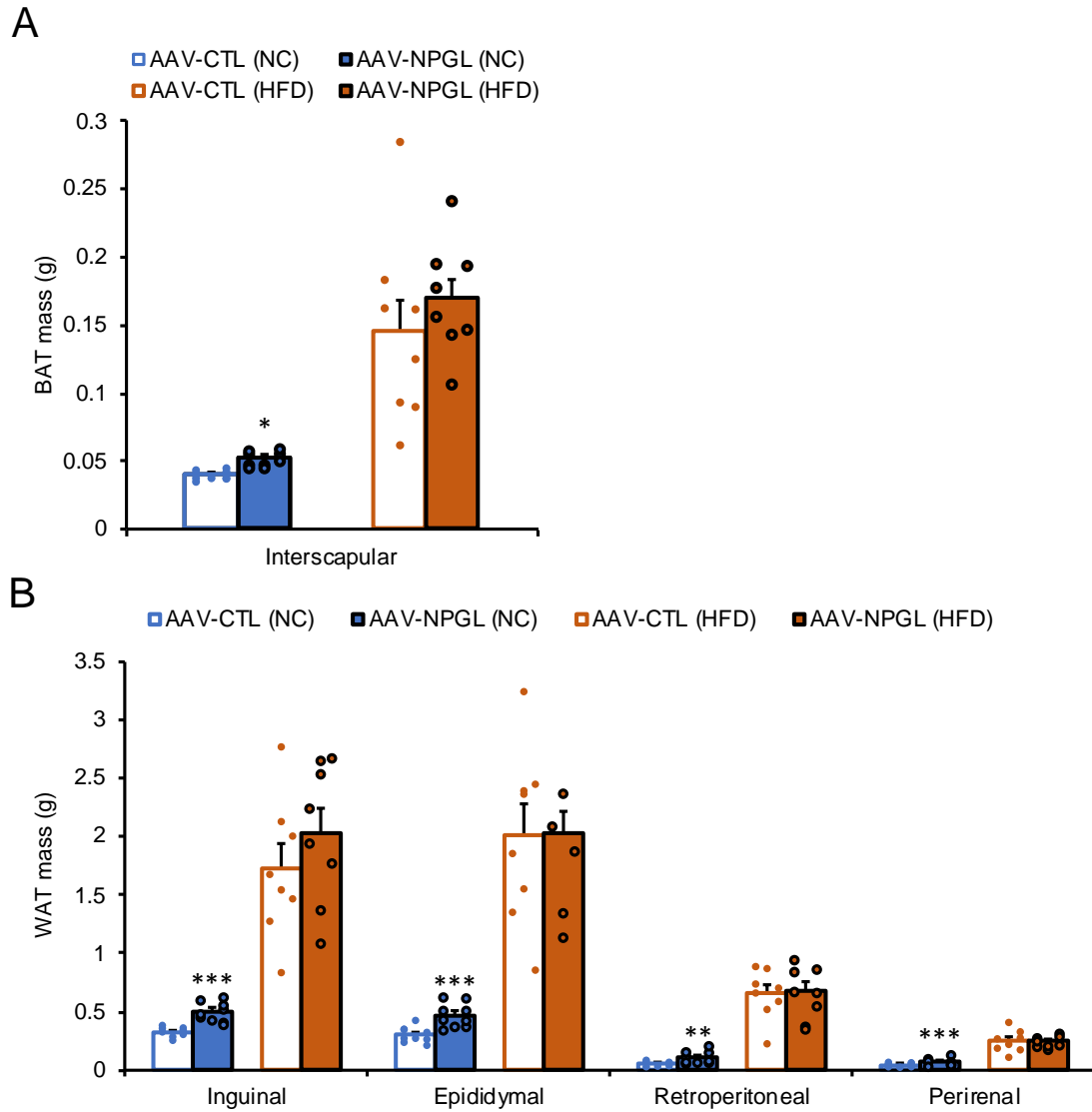


Figure S2. Effects of *Npgl* overexpression on fat accumulation in normal chow (NC)-fed and high-fat diet (HFD)-fed mice. The panels show the data obtained upon injection of the AAV-based control vector (AAV-CTL) or the AAV-based NPGL-precursor gene vector (AAV-NPGL) in NC-fed and HFD-fed mice. **(A)** Brown adipose tissue (BAT) mass. **(B)** Masses of inguinal, epididymal, retroperitoneal, and perirenal white adipose tissue (WAT). Circles represent individual data points. Each value represents the mean \pm standard error of the mean (NC-fed mice: $n = 9$; $*p < 0.05$, $**p < 0.01$, $***p < 0.005$ for Student's *t*-test, HFD-fed mice: $n = 8$).

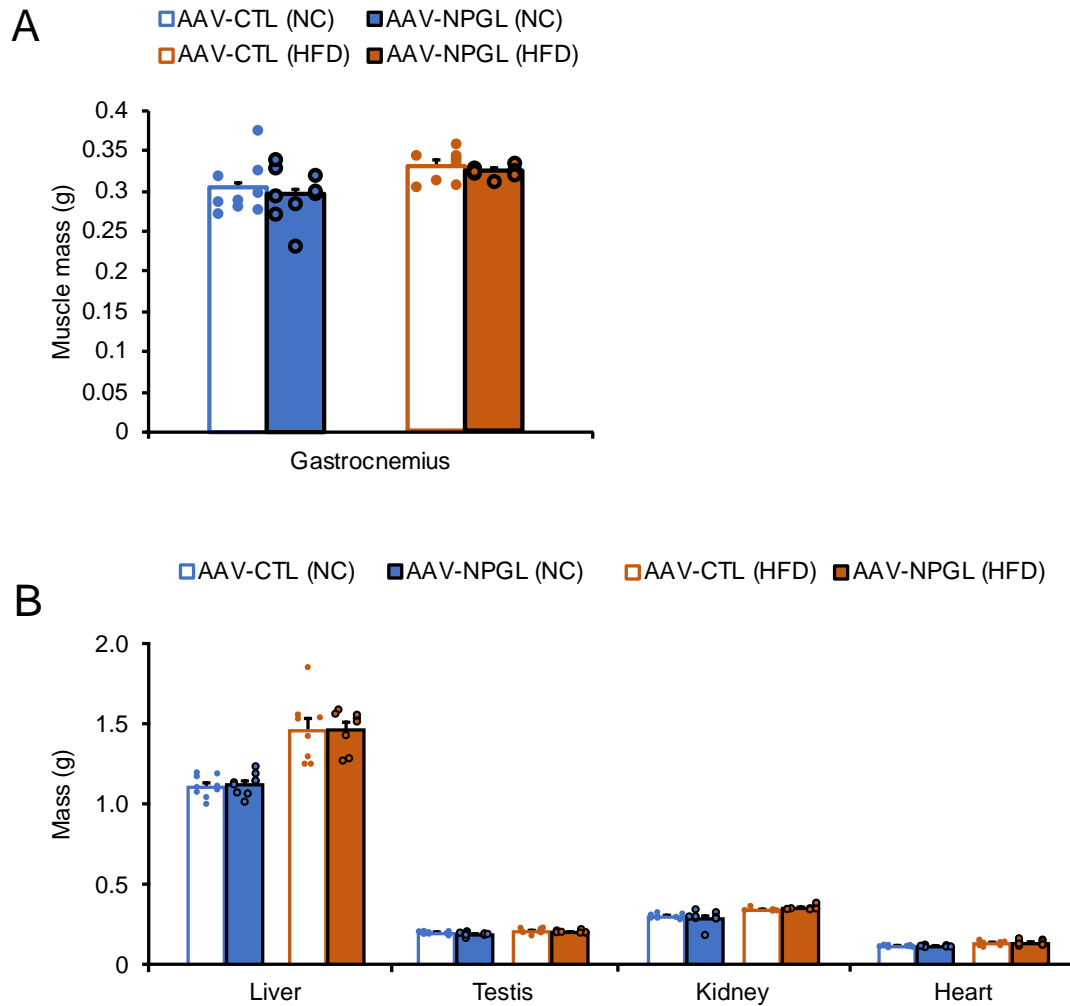


Figure S3. Effects of *Npgl* overexpression on muscle and organ masses in normal chow (NC)-fed and high-fat diet (HFD)-fed mice. The panels show the data obtained upon injection of the AAV-based control vector (AAV-CTL) or the AAV-based NPGL-precursor gene vector (AAV-NPGL) in NC-fed and HFD-fed mice. (A) Gastrocnemius muscle mass. (B) Masses of the liver, testis, kidney, and heart. Circles represent individual data points. Each value represents the mean \pm standard error of the mean (NC-fed mice: $n = 9$, HFD-fed mice: $n = 8$).

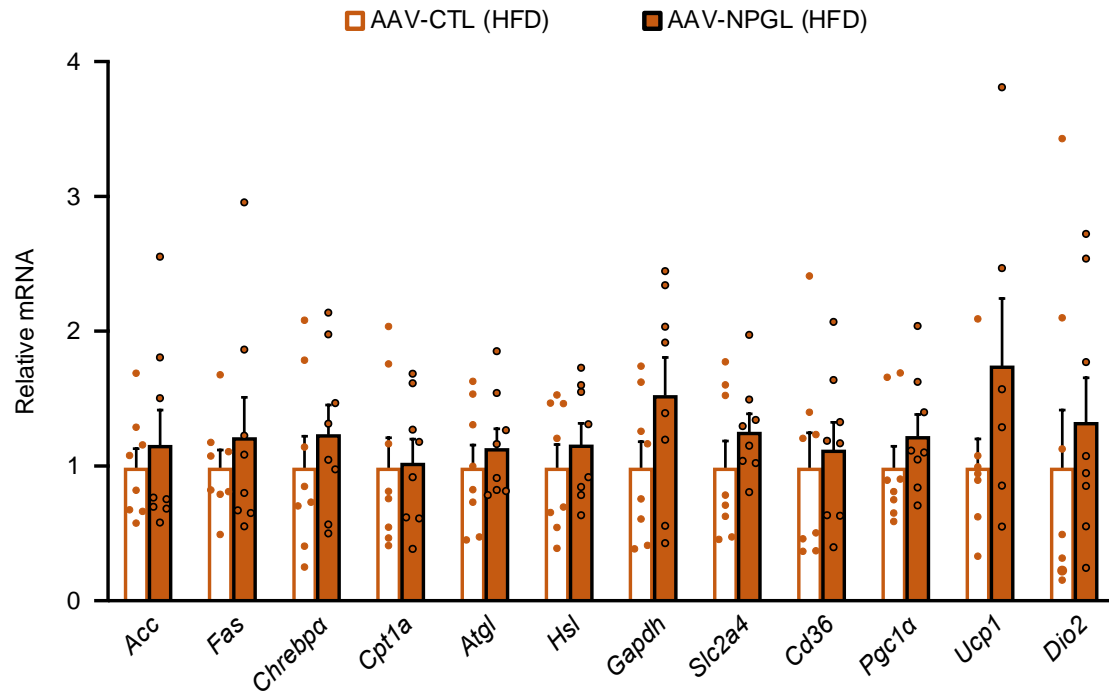


Figure S4. Effects of *Npgl* overexpression on mRNA expression of genes related to lipid metabolism, glycolysis, glucose and lipid uptake, and browning in the inguinal white adipose tissue of high-fat diet (HFD)-fed mice. The graph shows the data obtained upon injection of the AAV-based control vector (AAV-CTL) or the AAV-based NPGL-precursor gene vector (AAV-NPGL) in HFD-fed mice. mRNA expression levels of acetyl-CoA carboxylase (*Acc*), fatty acid synthase (*Fas*), carbohydrate-responsive element-binding protein α (*Chrebp* α), carnitine palmitoyltransferase 1a (*Cpt1a*), adipose triglyceride lipase (*Atgl*), hormone-sensitive lipase (*Hsl*), glyceraldehyde-3-phosphate dehydrogenase (*Gapdh*), solute carrier family 2 member 4 (*Slc2a4*), cluster of differentiation 36 (*Cd36*), peroxisome proliferator-activated receptor γ coactivator 1 α (*Pgc1* α), uncoupling protein 1 (*Ucp1*), and type II iodothyronine deiodinase (*Dio2*). Circles represent individual data points. Each value represents the mean \pm standard error of the mean ($n = 6-8$).

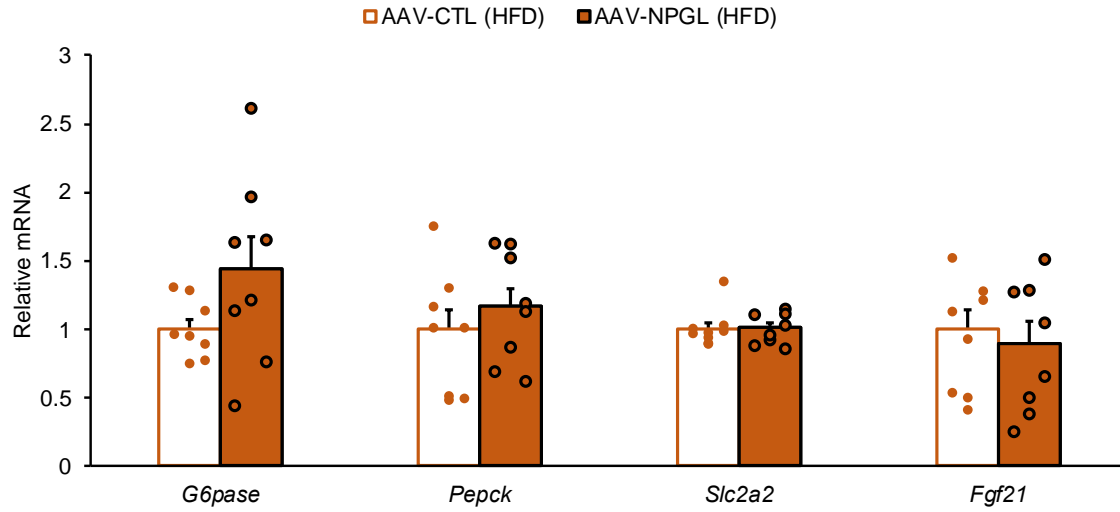


Figure S5. Effects of *Npgl* overexpression on mRNA expression of genes related to gluconeogenesis and glucose uptake in the liver of high-fat diet (HFD)-fed mice. The graph shows the data obtained upon injection of the AAV-based control vector (AAV-CTL) or the AAV-based NPGL-precursor gene vector (AAV-NPGL) in HFD-fed mice. mRNA expression levels of glucose-6-phosphatase (*G6pase*), phosphoenolpyruvate carboxykinase (*Pepck*), solute carrier family 2 member 2 (*Slc2a2*), and fibroblast growth factor 21 (*Fgf21*). Circles represent individual data points. Each value represents the mean \pm standard error of the mean ($n = 8$).