

Supplementary Fig. 1: Body weight, kidney weight, and kidney/body weight ratio of *Ptpn11*^{fl/fl} and *Ptpn11*^{fl/fl}; Pod-Cre mice treated (**a**) without (vehicle) and with STZ (160 μ g/g body weight, 20 weeks), and (**b**) fed chow and HFD (24 weeks). The sample number for each group was indicated in dot plots. * $p \le 0.05$, ** $p \le 0.01$ vehicle versus STZ (**a**), and chow versus HFD (**b**) of mice with the same genotype.



Supplementary Fig. 2: a,b Immunoblots of (**a**) pPERK T981, PERK, peIF2α S51, eIF2α, pNF-κB p65 S536, and NF-κB p65, and (**b**) mRNA expression of inflammatory cytokines IL1β, IL6, and TNFα in kidney samples (n=3 each group) from *Ptpn11^{fl/fl}* and *Ptpn11^{fl/fl}*; Pod-Cre mice fed chow and HFD (24 weeks). * $p \le 0.05$, ** $p \le 0.01$ vehicle versus STZ of mice with the same genotype. † $p \le 0.05$, †† $p \le 0.01$ *Ptpn11^{fl/fl}* versus *Ptpn11^{fl/fl}*; Pod-Cre under the same treatment. A.U.: arbitrary unit. c Confocal images of kidney sections from chow, STZ-treated (160 µg/g body weight, 20 weeks), and HFD-fed (24 weeks) *Ptpn11^{fl/fl}* and *Ptpn11^{fl/fl}*; and *Ptpn11^{fl/fl}*; Pod-Cre mice immunostained for Collagen III. Lower panel images are enlarged areas as highlighted by white boxes. Scale bar: 20 µm.



Supplementary Fig. 3: a Immunoblots of pNephrinY1176/Y1193, Nephrin, and GAPDH in kidney lysates from *Ptpn11^{fUfl}* and *Ptpn11^{fUfl}*; Pod-Cre mice treated without (vehicle) and with STZ (160 µg/g body weight, 20 weeks). Nephrin expression was normalized to GAPDH, and phosphorylation level was normalized to Nephrin. ** $p \le 0.01$ vehicle versus STZ of mice with the same genotype. † $p \le 0.05$, †† $p \le 0.01$ *Ptpn11^{fUfl}* versus *Ptpn11^{fUfl}*; Pod-Cre under the same treatment. A.U.: arbitrary unit. **b** Confocal images of kidney sections from chow, STZ-treated (160 µg/g body weight, 20 weeks), and HFD-fed (24 weeks) *Ptpn11^{fUfl}* and *Ptpn11^{fUfl}*; Pod-Cre mice immunostained for Nephrin. Lower panel images are enlarged areas as highlighted by white boxes. Scale bar: 20 µm.