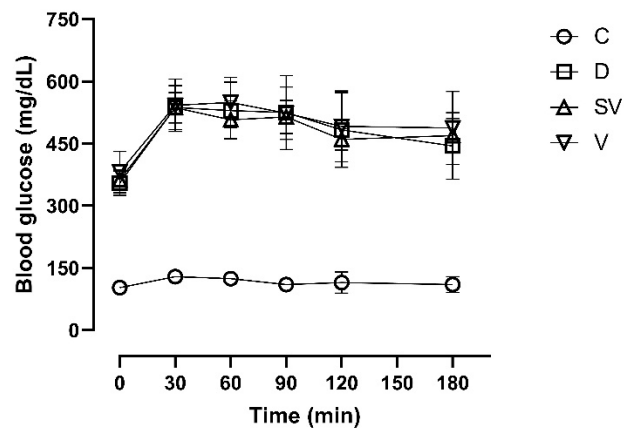


Supplementary Figure S1. Blood glucose levels during treatment. **(a)** Weekly measured blood glucose levels starting after treatment (means \pm SD). **(b)** Blood glucose change. C, Control (n=13); D, Diabetic (n=17); SV, Sacubitril/valsartan treated diabetic (n=16); V, Valsartan treated diabetic (n=15); each data point is one animal.



Supplementary Figure S2. Time course blood glucose levels during OGTT. C, Control (n=8); D, Diabetic (n=7); SV, Sacubitril/valsartan treated diabetic (n=8); V, Valsartan treated diabetic (n=9); values show as means \pm SD.

Supplementary Table S1. Blood glucose levels at each time point during OGTT

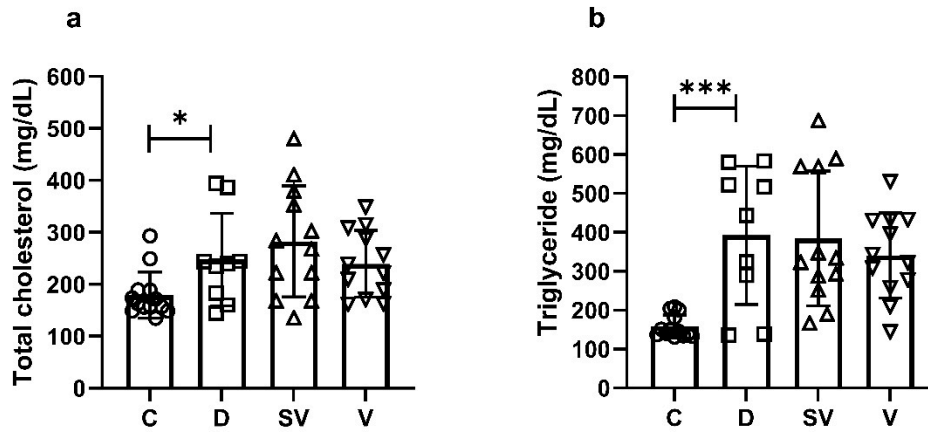
C, Control (n=8); D, Diabetic (n=7); SV, Sacubitril/valsartan treated diabetic (n=8); V, V, Valsartan treated diabetic (n=9). ****, $p < 0.0001$ compared to control. Values shown as means \pm SD.

	C (n=8)	D (n=7)	SV (n=8)	V (n=9)
0. min BG (mg/dL)	102.38 \pm 7.95	354.71 \pm 29.25****	362.13 \pm 24.19	381.33 \pm 50.14
30. min BG (mg/dL)	128.88 \pm 10.36	536.86 \pm 53.06****	536.13 \pm 36.72	542.00 \pm 63.31
60. min BG (mg/dL)	124.13 \pm 13.16	529.14 \pm 68.54****	507.25 \pm 45.60	549.11 \pm 60.58
90. min BG (mg/dL)	110.13 \pm 12.81	524.86 \pm 89.11****	513.88 \pm 40.02	523.00 \pm 63.04
120. min BG (mg/dL)	114.63 \pm 25.49	482.57 \pm 90.48****	460.13 \pm 25.96	491.22 \pm 85.40
180. min BG (mg/dL)	109.88 \pm 18.94	444.00 \pm 79.86****	469.13 \pm 41.35	487.33 \pm 88.40

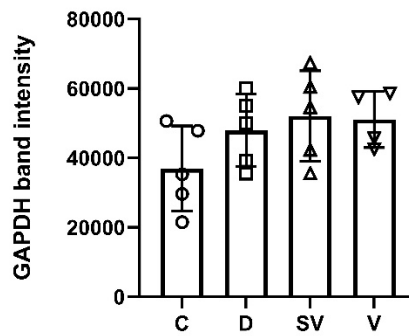
Supplementary Table S2. Fasting blood glucose and insulin levels prior to OGTT and HOMA-IR values

C, Control (n=5); D, Diabetic (n=5); SV, Sacubitril/valsartan treated diabetic (n=6); V, Valsartan treated diabetic (n=5). ****, $p < 0.0001$ compared to control. Values shown as means \pm SD,

	C (n=5)	D (n=5)	SV (n=6)	V (n=5)
Fasting blood glucose (mg/dL)	102.00 \pm 8.34	350.40 \pm 27.49****	365.00 \pm 24.39	400.20 \pm 62.54
Fasting insulin (mIU/L)	10.21 \pm 0.77	10.77 \pm 0.96	10.30 \pm 0.86	10.36 \pm 0.63
HOMA-IR	2.56 \pm 0.20	9.27 \pm 0.45****	9.29 \pm 1.18	10.22 \pm 1.74



Supplementary Figure S3. Plasma lipid levels. **(a)** Total cholesterol levels. **(b)** Triglyceride levels. C, Control (n=13); D, Diabetic (n=9); SV, Sacubitril/valsartan treated diabetic (n=12); V, Valsartan treated diabetic (n=12). *, $p < 0.05$; ***, $p < 0.001$ compared to control. Each data point represents one animal.



Supplementary Figure S4. GAPDH validation as a housekeeping gene. C, Control (n=5); D, Diabetic (n=5); SV, Sacubitril/valsartan treated diabetic (n=5); V, Valsartan treated diabetic (n=4). Each datapoint represents one animal.