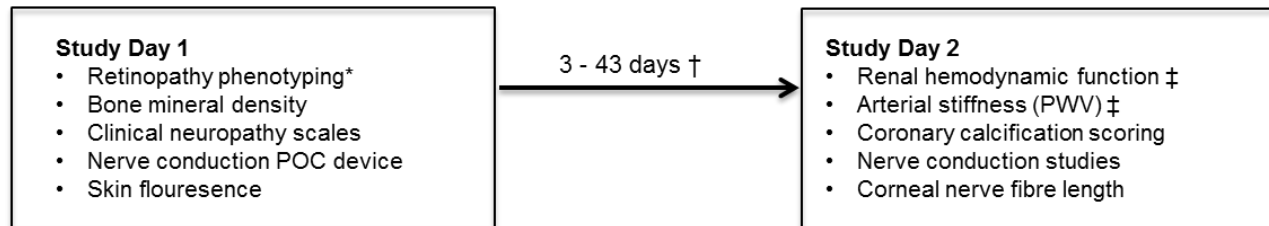


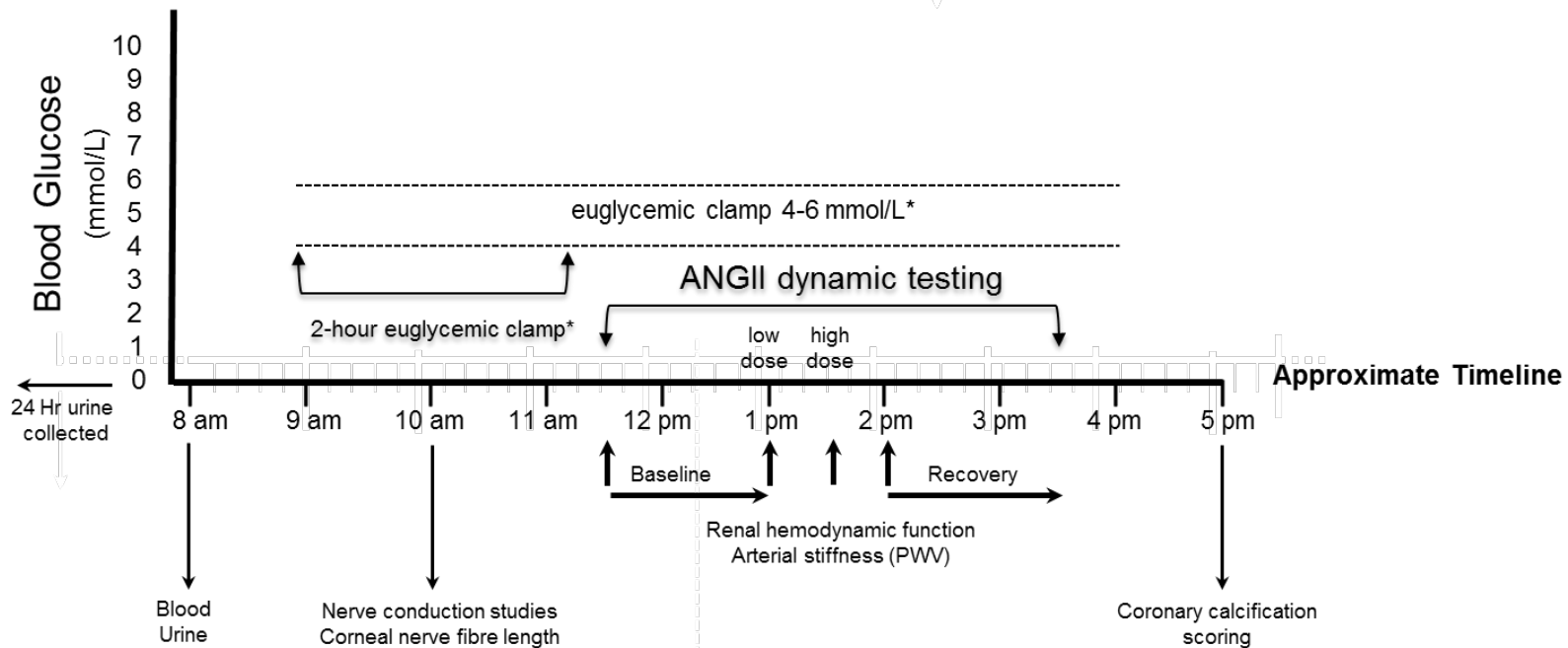
SUPPLEMENTAL DATA

Supplemental Figure 1. Study design and experimental protocol

A) Study Design



B) Experimental Protocol



Legend: *Type 1 diabetes only. † RAASi withdrawal. ‡ At baseline and during ANGII

ANGII, angiotensin II; POC, point of care; PWV, pulse wave velocity; RAASi, renin-angiotensin-aldosterone system inhibitor.

Supplemental Table 1. Heart rate variability correlations in DKD resistors (A) and DKD (B) in type 1 diabetes in response to exogenous RAAS stimulation with ANGII

(A) DKD resistors

	Δ RVR in response to low dose ANG II	Δ RVR in response to high dose ANGII	Δ SBP in response to low dose ANGII	Δ SBP in response to high dose ANGII
RMSSD*	r=0.41, p=0.009	r=0.36, p=0.021	r=0.28, p=0.075	r=0.14, p=0.39
SDNN*	r=0.31, p=0.051	r=0.27, p=0.10	r=0.23, p=0.14	r=0.08, p=0.63
SDNN/RMSSD	r=-0.40, p=0.010	r=-0.40, p=0.012	r=-0.19, p=0.22	r=-0.12, p=0.46
LF/HF Ratio*	r=-0.40, p=0.012	r=-0.48, p=0.002	r=-0.07, p=0.65	r=-0.22, p=0.16

* Natural log-transformed

(B) DKD

	Δ RVR in response to low dose ANG II	Δ RVR in response to high dose ANG II	Δ SBP in response to low dose ANGII	Δ SBP in response to high dose ANG II
RMSSD*	r=0.17, p=0.50	r=-0.03, p=0.89	r=0.47, p=0.035	r=0.14, p=0.57
SDNN*	r=0.04, p=0.86	r=-0.25, p=0.31	r=0.29, p=0.22	r=-0.04, p=0.88
SDNN/RMSSD	r=-0.26, p=0.27	r=-0.15, p=0.55	r=-0.43, p=0.053	r=-0.24, p=0.33
LF/HF Ratio*	r=-0.29, p=0.22	r=-0.34, p=0.16	r=-0.43, p=0.060	r=-0.29, p=0.23

* Natural log-transformed

DKD, diabetic kidney disease; ANGII, angiotensin II; HF, high frequency; LF, low frequency; RMSSD, root mean square successive difference; SDNN, sympathetic activity, standard deviation of normal-to-normal interval; RVR, renal vascular resistance.