

**S2 Table. Uni-nephrectomised db/db mice, general measures**

		lean vehicle (n=9)	obese w18 (n=10)	obese vehicle (n=9)	obese AZD9977 (n=8)	obese eplerenone (n=9)	obese enalapril (n=9)	obese enalapril + AZD9977 (n=7)	obese enalapril + eplerenone (n=9)
<b>BW (g)</b>	w18	27.9 ± 0.5	49.7 ± 2.5	50.3 ± 2.2	48 ± 2.3	49.1 ± 2	47.5 ± 1.5	50.2 ± 2.2	49.4 ± 2.1
	w22	29.1 ± 0.5*		46.6 ± 2.5	41.4 ± 1.9	42.5 ± 1.8	39.3 ± 1.8	39.9 ± 2.3	38.3 ± 3.5*
<b>BG (mM)</b>	w18	8.7 ± 0.5	>29.4	>29.4	>29.4	>29.4	>29.4	>29.4	>29.4
	w22	9.1 ± 0.5		>29.4	>29.4	>29.4	>29.4	>29.4	>29.4
<b>HbA1c (%)</b>	w18	4.7 ± 0.1	>12	>12	>12	>12	>12	>12	>12
	w22	4.7 ± 0.1		>12	>12	>12	>12	>12	>12
<b>UAE (µg 24h<sup>-1</sup>)</b>	w18	3 ± 1	867 ± 155	907 ± 138	797 ± 169	892 ± 186	958 ± 184	1055 ± 296	832 ± 129
	w22	1 ± 0		1023 ± 175	588 ± 80 *	569 ± 76 *	583 ± 97 *	356 ± 52 *	244 ± 51 *
<b>u-Na<sup>+</sup>/K<sup>+</sup></b>	end		0.53 ± 0.01*	0.39 ± 0.01	0.47 ± 0.01*	0.48 ± 0.02*	0.40 ± 0.01	0.47 ± 0.01*	0.50 ± 0.02*
<b>u-Na<sup>+</sup> (mmol/24h)</b>	end		0.52 ± 0.04*	0.37 ± 0.02	0.45 ± 0.05	0.40 ± 0.05	0.36 ± 0.03	0.39 ± 0.03	0.39 ± 0.04
<b>u-K<sup>+</sup> (mmol/24h)</b>	end		0.98 ± 0.06	0.94 ± 0.04	0.97 ± 0.10	0.85 ± 0.10	0.90 ± 0.07	0.84 ± 0.07	0.80 ± 0.10
<b>p-K<sup>+</sup> (mM)</b>	end	3.2 ± 0.1		3.6 ± 0.1	3.9 ± 0.2	3.8 ± 0.1	4.4 ± 0.1*	4.8 ± 0.2*	4.7 ± 0.3*
<b>FE K<sup>+</sup> (%)</b>	end			20.5 ± 1.2 <sup>#</sup>	20.1 ± 0.9 <sup>#</sup>	22.3 ± 1.0 <sup>#</sup>	15.6 ± 0.4*	15.6 ± 0.8*	17.8 ± 1.0
<b>FE Na<sup>+</sup> (%)</b>	end			0.19 ± 0.01	0.24 ± 0.01	0.27 ± 0.02*	0.18 ± 0.01	0.24 ± 0.02	0.27 ± 0.02*
<b>p-crea (µg mL<sup>-1</sup>)</b>	end	9.9 ± 0.5*	5.1 ± 0.2	5.3 ± 0.3	5.8 ± 0.2	5.9 ± 0.2	5.0 ± 0.2	5.6 ± 0.3	6.4 ± 0.6
<b>C<sub>drug</sub> (µM)</b>					0.35 ± 0.04	0.15 ± 0.02		0.27 ± 0.05	0.12 ± 0.03
<b>Fold IC<sub>50</sub></b>					2.4	1.1		1.9	0.9

BW: body weight; BG: blood glucose; UAE: urinary albumine excretion; u-Na<sup>+</sup>/K<sup>+</sup>: urine Na<sup>+</sup>/K<sup>+</sup> ratio; u-Na<sup>+</sup>: amount Na<sup>+</sup> excreted in urine over 24h; u-K<sup>+</sup>: amount K<sup>+</sup> excreted in urine over 24h; p-K<sup>+</sup>: plasma K<sup>+</sup>; FE K<sup>+</sup>: fractional K<sup>+</sup> excretion; FE Na<sup>+</sup>: fractional Na<sup>+</sup> excretion; p-crea: plasma creatinine; C<sub>drug</sub>: terminal drug exposure levels. Unbound fraction in plasma (AZD9977 55%, eplerenone 69%) was used to convert C<sub>drug</sub> to free plasma levels; Fold IC<sub>50</sub>: free plasma levels related to *in vitro* potencies (free plasma levels / *in vitro* IC<sub>50</sub>). Average ± SEM. \*p<0.05 vs obese vehicle at w22, #p<0.05 vs enalapril.