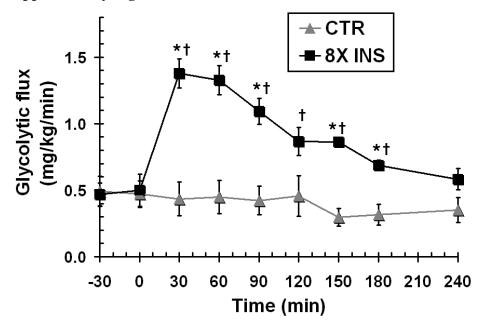
**Supplementary Table 1**. Arterial blood/plasma levels, net hepatic uptake, and net hepatic fractional extraction of selected gluconeogenic substrates in the 24 h fasted conscious dog subjected to either euinsulinemia (Control) or hyperinsulinemia (8X INS). Somatostatin, insulin and glucagon were infused from 0-min to control hormone levels and glucose was infused peripherally to maintain euglycemia.

	Basal	Experimental Period (min)					
	Period	30	60	90	120	180	240
Arterial Plasma Glycerol Level (µmol/L)							
Control	$97 \pm 9$	$86 \pm 7$	84 ± <u>9</u>	$87 \pm 13$	$79 \pm 10$	$77 \pm 4$	$80 \pm 11$
8X INS	$97 \pm 5$	$48 \pm 5*\dagger$	$39 \pm 5*†$	$32 \pm 2*\dagger$	$30 \pm 2*\dagger$	25 ± 2*†	$35 \pm 4*†$
Net Hepatic Glycerol Uptake (µmol/kg/min)							
Control	1.9 ±0.2	1.6 ±0.2	$1.6 \pm 0.1$	$1.6 \pm 0.1$	$2.0 \pm 0.3$	1.6 ±0.1	1.9 ±0.2
8X INS	2.1 ±0.2	0.8 ±0.1*†	0.7 ±0.1*†	0.5 ±0.1*†	0.6 ±0.1*†	0.5 ±0.1*†	0.8 ±0.1*†
Arterial Blood Alanine Level (µmol/L)							
Control	391 ±82	405 ±77	428 ±78	453 ±78	467 ±64	443 ±52	438 ±63
8X INS	329 ±31	324 ±31	326 ±35	270 ±24*	196 ±8*†	180 ±5*†	152 ±6*†
Net Hepatic Alanine Uptake (µmol/kg/min)							
Control	$3.3 \pm 0.7$	$3.5 \pm 0.6$	$3.3 \pm 0.7$	$4.2 \pm 0.7$	$3.9 \pm 0.9$	$3.8 \pm 1.0$	$3.5 \pm 0.6$
8X INS	$3.1 \pm 0.3$	$2.9 \pm 0.3$	$3.1 \pm 0.6$	$3.6 \pm 0.6$	$3.4 \pm 0.4$	$3.0 \pm 0.1$	$2.9 \pm 0.1$
Net Hepatic Alanine Fractional Extraction							
Control	0.26±0.05	0.25±0.08	0.22±0.08	0.29±0.05	$0.28 \pm 0.05$	$0.29 \pm 0.05$	$0.30 \pm 0.04$
8X INS	$0.25 \pm 0.03$	$0.24 \pm 0.02$	$0.25 \pm 0.01$	0.34 ±.03†	0.39 ±0.01*†	0.41 ±0.02*†	0.45 ±0.01*†
Arterial Blood GNG Amino Acid Level (µmol/L)							
Control	$1722 \pm 189$	$1819 \pm 187$	1861 ± 188	$1913 \pm 213$	$1914 \pm 213$	$2028 \pm 231$	$2021 \pm 350$
8X INS	1619 ±96	1527 ±95	1465 ±76	1359 ±68*	1252 ±70*†	1143 ±42*†	1096 ±38*†
Net Hepatic GNG Amino Acid Uptake (µmol/kg/min)							
Control	$4.4 \pm 1.2$	$5.1 \pm 2.0$	$4.0 \pm 1.4$	$5.4 \pm 0.9$	$5.1 \pm 1.3$	$5.4 \pm 1.1$	$5.6 \pm 0.3$
8X INS	$4.6 \pm 0.5$	$4.1 \pm 0.5$	$4.9 \pm 0.6$	$5.7 \pm 0.5$	$5.5 \pm 0.5$	$4.9 \pm 0.5$	$5.6 \pm 0.4$
Net Hepatic GNG Amino Acid Fractional Extraction							
Control	$0.10 \pm 0.02$	$0.12 \pm 0.04$	$0.08 \pm 0.03$	0.12 ±0.03	$0.12 \pm 0.03$	$0.13 \pm 0.05$	$0.10 \pm 0.02$
8X INS	0.09 ±0.01	$0.12 \pm 0.01$	0.15 ±0.01*†	0.15 ±0.02†	0.16 ±0.01†	$0.13 \pm 0.01 \dagger$	0.14 ±0.01*†

Mean  $\pm$  S.E.M; n=6 and n=30 in Control and 8X INS groups, respectively. \*P<0.05 vs. Control group, † P<0.05 vs. basal period.

## **Supplementary Figure 1**



Supplemental Figure 1. Glycolytic flux in 24 h fasted conscious dogs during the basal (-30 to 0 min) and experimental (0-240 min) periods. No persisting alteration in GNG flux-to-G6P was observed in either group. Thus, calculation for glycolytic flux assumed GNG flux-to-G6P was constant at basal rates and that lactate flux was unidirectional in the control period. Data are means  $\pm$  S.E.M, n=7 in CTR and n=20 in 8X INS groups. \*P<0.05 vs. CTR group; †P<0.05 vs. basal period.