

SUPPLEMENTARY DATA

**Supplementary Table 1.** Liver ATP levels phosphorylated AMPK levels (as previously described (29,26)), GFAT mRNA levels (Mm00600127\_m1), FASN mRNA levels (Mm00662319\_m1), SCD1 mRNA levels (Mm01197142\_m1), SREBF1 levels (Mm00550338\_m1) (with 18s rNA as control) and plasma adiponectin concentration (Millipore RIA) in transgenic mice and negative littermates, approximately 16 weeks of age (n = 4-5 for each group).

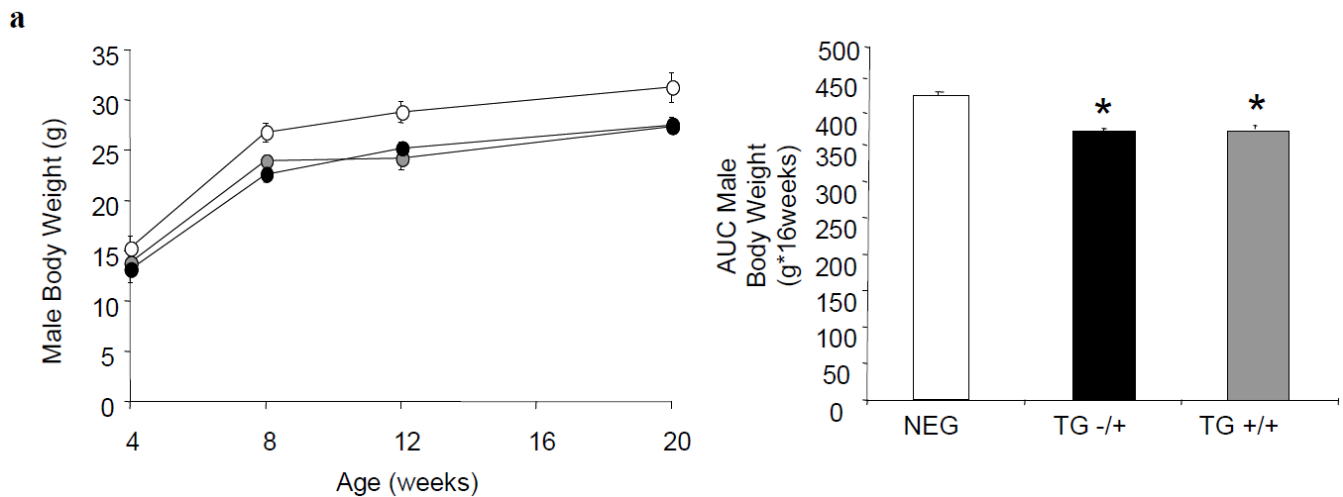
	NEG	TG
ATP ( $\mu\text{g}/\text{mg}$ protein)	$3.8 \pm 1.1$	$4.9 \pm 0.8$
pAMPK (Fold of NEG)	$1.0 \pm 0.3$	$0.8 \pm 0.2$
GFAT mRNA (Fold of NEG)	$1.00 \pm 0.13$	$1.40 \pm 0.15$
FASN mRNA (Fold of NEG)	$1.00 \pm 0.34$	$0.93 \pm 0.23$
SCD1 mRNA (Fold NEG)	$1.00 \pm 0.22$	$1.08 \pm 0.39$
SREBF1 mRNA (Fold NEG)	$1.00 \pm 0.21$	$0.94 \pm 0.07$
Plasma Adiponectin ( $\mu\text{g}/\text{mL}$ )	$16.9 \pm 1.7$	$15.6 \pm 1.8$

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**Supplementary Table 2.** Hepatic fructose-1,6-bisphosphate and fructose-6-phosphate. Liver homogenates were used to measure fructose-1,6-bisphosphate and fructose-6-phosphate as previously described (29). Fasted levels of gluconeogenic intermediates from transgenic and negative littermates, ~24 weeks of age (n = 3-4 for each group, \*p<0.05 vs. NEG).

	NEG	TG
Fructose-1,6-bisphosphate (mmol/min/μg protein)	0.11 ± 0.01	*0.43 ± 0.07
Fructose-6-phosphate (mmol/min/μg protein)	0.99 ± 0.46	*3.84 ± 0.90

**Supplementary Figure 1.** Characterisation of hepatic male FBPase transgenic mice. Body weights of negative (○), hemizygous transgenic (●) and homozygous transgenic (◐) male mice from homozygous breeding (a) (n=7-10, AUC \*p<0.05 vs. NEG).



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**Supplementary Figure 2.** Characterisation of transgenic mice over-expressing FBPase in the liver and hypothalamus. Body weights of male negative (○) and transgenic (●) mice from hemizygous breeding from 4 weeks of age to 22 weeks old (a) (n=10, AUC \*p<0.05 vs. NEG). Body weights of female negative (○) and transgenic (●) mice from hemizygous breeding (b) (n=10, AUC \*p<0.05 vs. NEG). Fat pad mass of both female and male hemizygous transgenic mice (black bars) compared with their negative littermates at 24 weeks of age (white bars) (c) (n=10 for NEG, \*p<0.05). Average daily food intake of both female and male hemizygous transgenic mice (black bars) compared with negative littermates (white bars) (d) (n=10, \*p<0.05 vs. NEG). Physical activity measurements at 23 weeks of age of female and male negative (white bars) and hemizygous transgenic mice (black bars) (e) (n=10).

