

## SUPPLEMENTARY DATA

**Figure S1. Tissue specific *Vhl* inactivation following Ad-Cre tail vein injection leads to fatty liver.** (A) PCR analysis of genomic DNA isolated from tissues of *Vhl*<sup>+/+</sup> and *Vhl*<sup>F/F</sup> animals at the indicated number of days post Ad-Cre injection; 1:1 mixture of unrecombined (*Vhl*<sup>F</sup>) and efficiently recombined (*Vhl*<sup>Frec</sup>) genomic DNA shown as a control. (B) Microscopic analysis of representative liver sections of animals of the indicated genotypes after Ad-Cre injection. H&E, hematoxylin eosin staining; ORO, Oil red O stained cryosections counterstained with hematoxylin; scale bar, 50μm.

**Table S1**

Carnitine and acylcarnitine levels in livers of nonfasted and 24 hours fasted animals sacrificed on day 4 after Ad-Cre injection. Values are mean  $\pm$  standard deviation and are presented as nmol/g of liver . \*  $p < 0.05$ , \*\*  $p < 0.01$  when compared to wild-type under the same conditions.

<b>Nonfasted</b>	<b><i>Vhl</i><sup>+/+</sup> (n = 7)</b>	<b><i>Vhl</i><sup>F/F</sup> (n = 7)</b>
Free carnitine	415.4 $\pm$ 83.4	306.8 $\pm$ 39.9**
Acetylcarnitine (C2)	18.4 $\pm$ 22.2	56 $\pm$ 25.1*
Propionylcarnitine (C3)	5.3 $\pm$ 3.2	6.1 $\pm$ 1.8
Butyrylcarnitine (C4)	0.83 $\pm$ 0.59	2.71 $\pm$ 1.38**
Isovalerylcarnitine (C5)	0.86 $\pm$ 0.2	0.61 $\pm$ 0.15*
Octanoylcarnitine (C8)	0.0019 $\pm$ 0.0005	0.0029 $\pm$ 0.0013
Myristoylcarnitine (C14)	0.0025 $\pm$ 0.0012	0.0123 $\pm$ 0.0162
Palmitoylcarnitine (C16)	0.044 $\pm$ 0.031	0.192 $\pm$ 0.238
<b>Fasted</b>	<b><i>Vhl</i><sup>+/+</sup> (n = 6)</b>	<b><i>Vhl</i><sup>F/F</sup> (n = 7)</b>
Free carnitine	622.3 $\pm$ 196.5	366.1 $\pm$ 51.7**
Acetylcarnitine (C2)	10.1 $\pm$ 5	26.2 $\pm$ 15.5*
Propionylcarnitine (C3)	3.4 $\pm$ 1.8	3.8 $\pm$ 1.1
Butyrylcarnitine (C4)	0.61 $\pm$ 0.28	1.49 $\pm$ 0.85*
Isovalerylcarnitine (C5)	0.85 $\pm$ 0.45	0.5 $\pm$ 0.15
Octanoylcarnitine (C8)	0.0017 $\pm$ 0.0007	0.0035 $\pm$ 0.0014*
Myristoylcarnitine (C14)	0.0168 $\pm$ 0.007	0.0139 $\pm$ 0.0068
Palmitoylcarnitine (C16)	0.146 $\pm$ 0.117	0.156 $\pm$ 0.097

## Table S2

Sequences of primers used in PCR analysis.

Primer name	Sequence 5' → 3'
Vhl F1	TGCCTGGTACCCACGAAAGTGTC
Vhl R1	CTGACTTCCACTGATGCTTGTCACAG
Vhl F2	CCGGAGTAGGATAAGTCAGCTGAG
Hif1b F2	TGCCAACATGTGCCACCATGT
Hif1b R2	GTGAGGCAGATTTCTTCCATGCTC
Hif1b F4	ACGCACTACAACACCTGAGCTAA

**Table S3**

Sequences of primers used in qRT-PCR analysis.

Protein	Gene (MGI)	Sequence 5'-3'	Reference/Note
Cyclophilin b	<i>Ppib</i>	TGGAGAGCACCAAGACAGACA TGCCGAGTCGACAATGAT	[1]
Vascular endothelial growth factor	<i>Vegf</i>	CACGACAGAAGGAGAGCAGAA CGCTGGTAGACGTCCATGA	Nguyen et al., unpublished
Glucose transporter 1	<i>Glut1</i>	GGTGTGCAGCAGCCTGTGTA CAACAAACAGCGACACCACAGT	
Glucose transporter 1	<i>Glut1</i>	CCTGCTCATCAATCGTAACGAGG CGACCCTCTTCTTTCATCTCC	
SREBP cleavage-activating protein	<i>SCAP</i>	ATTGCTCACCGTGGAGATGTT GAAGTCATCCAGGCCACTACTAATG	[2]
Insulin induced gene 1	<i>Insig-1</i>	TCACAGTGACTGAGCTTCAGCA TCATCTTCATCACACCAGGAC	[3]
Insulin induced gene 2a	<i>Insig-2a</i>	CCCTCAATGAATGTACTGAAGGATT TGTGAAGTGAAGCAGACCAATGT	[3]
Insulin induced gene 2b	<i>Insig-2b</i>	CCGGGCAGAGCTCAGGAT GAAGCAGACCAATGTTCAATGG	[3]
Sterol regulatory element binding protein 1a	<i>SREBP-1a</i>	GGCCGAGATGTGCGAACT TTGTTGATGAGCTGGAGCATGT	[2]
Sterol regulatory element binding protein 1c	<i>SREBP-1c</i>	GGAGCCATGGATTGCACATT GGCCCGGAAGTCACTGT	[2]
Sterol regulatory element binding protein 2	<i>SREBP-2</i>	GCGTCTGGAGACCATGGA ACAAAGTTGCTCTGAAAACAAATCA	[2]
Farnesyl diphosphate synthase	<i>Fpps</i>	ATGGAGATGGGCGAGTTCCTC CCGACCTTCCCGTCACA	[2]
Squalene Synthase, Farnesyl diphosphate farnesyl transferase 1	<i>Fdfl</i>	CCAACTCAATGGGTCTGTTCCT TGGCTTAGCAAAGTCTTCCAAC	[2]
3-hydroxy-3-methylglutaryl-Coenzyme A synthase 1	<i>Hmgcs1</i>	GCCGTGAACTGGGTCGAA GCATATATAGCAATGTCTCCTGCAA	[2]
3-hydroxy-3-methylglutaryl-Coenzyme A synthase 2 (mitochondrial)	<i>Hmgcs2</i>	GACTTCCTGTCATCCAGC GGTGTAGGTTTCTTCCAGC	
3-hydroxy-3-methylglutaryl-Coenzyme A reductase	<i>Hmgcr</i>	CTGTGGAATGCCTTGTGATTG AGCCGAAGCAGCACATGAT	[2]
Low density lipoprotein receptor	<i>Ldlr</i>	AGGCTGTGGGCTCCATAGG TGCGGTCCAGGGTCATCT	[2]
Very low density lipoprotein receptor	<i>Vldlr</i>	ACCTGTTCTGTCCCAATGG TCACTGTAAGTCACAGGAGTTGAAGTAC	

Peroxisome proliferator-activated receptor $\alpha$	<i>Ppara</i>	ACAAGGCCTCAGGGTACCA GCCGAAAGAAGCCCTTACAG	[4]
Peroxisome proliferator-activated receptor $\alpha$	<i>Ppara</i>	G TTCACGCATGTGAAGGCTGTAAG CATTGTGTGACATCCCGACAGAC	
Peroxisome proliferator-activated receptor $\delta$	<i>Ppard</i>	ACGCACCCTTTGTCATCCA TTCCACACCAGGCCCTTCT	
Peroxisome proliferator-activated receptor $\gamma$	<i>Pparg</i>	CACAATGCCATCAGGTTTGG GCTGGTCGATATCACTGGAGATC	[4]
Liver X receptor $\alpha$	<i>Nr1h3</i>	TCTGGAGACGTCACGGAGGTA CCCGTTGTAAGTGAAGTCCTT	[5]
Retinoid X receptor $\alpha$	<i>Rxra</i>	CCTTACCAAGCACATCTG TG TAGGTCAGGTCTTTGCG	
ATP-binding cassette protein A1	<i>Abca1</i>	CGTTTCCGGAAGTGCCTA GCTAGAGATGACAAGGAGGATGGA	[1]
ATP-binding cassette protein G5	<i>Abcg5</i>	TGGATCCAACACCTCTATGCTAAA GGCAGGTTTTCTCGATGAACTG	[1]
Cholesterol 7-a hydroxylase; Cytochrome P450, family 7, subfamily A, polypeptide 1	<i>Cyp7a1</i>	AGCAACTAAACAACCTGCCAGTACTA GTCCGGATATTCAAGGATGCA	[1]
Malic enzyme	<i>Mel1</i>	GCCGGCTCTATCCTCCTTTG TTTGTATGCATCTTGACAATCTTT	[1]
Acyl-CoA synthetase short-chain family member 2	<i>Acss2</i>	GCTGCCGACGGGATCAG TCCAGACACATTGAGCATGTCAT	[1]
Acetyl-CoA carboxylase 1	<i>Acaca</i>	TGGACAGACTGATCGCAGAGAAAAG TGGAGAGCCCCACACACA	[2]
Fatty Acid Synthase	<i>Fasn</i>	GCTGCGGAAACTTCAGGAAAT AGAGACGTGTCACCTCCTGGACTT	[2]
Stearoyl CoA Desaturase 1	<i>Scd1</i>	CCGAGACCCCTTAGATCGA TAGCCTGTAAGATTTCTGCAAACC	[2]
Glycerol-3-phosphate acyltransferase, mitochondrial	<i>Gpam</i>	CAACACCATCCCCGACATC GTGACCTTCGATTATGCGATCA	[1]
Diacylglycerol acyltransferase 1	<i>Dgat1</i>	GAGGCCTCTGCCCCTATG GCCCTGGACAACACAGACT	[6]
Diacylglycerol acyltransferase 2	<i>Dgat2</i>	CCGCAAAGGCTTTGTGAAG GGAATAAGTGGGAACCAGATCA	[6]
Acyl glycerol-3-phosphate acyltransferase 1	<i>Agpat1</i>	GCTGGCTGGCAGGAATCAT GTCTGAGCCACCTCGGACAT	[6]
Acyl glycerol-3-phosphate acyltransferase 2	<i>Agpat2</i>	TTTGAGGTCAGCGGACAGAA AGGATGCTCTGGTGATTAGAGATGA	[6]
Acyl glycerol-3-phosphate acyltransferase 3	<i>Agpat3</i>	CCAGTGGCTTCACAAGCTGTAC CCCTGGGAATACACCCTTCTG	
Acyl glycerol-3-phosphate acyltransferase T4	<i>Agpat4</i>	ACTTCGTGGAAATGATCTTTTGC GAGGTGCAGCAGGCTCTTG	

Acyl-Coenzyme A oxidase 1, palmitoyl	<i>Acox1</i>	AGATTGGTAGAAATTGCTGCAAAA ACGCCACTTCCTTGCTCTTC	
Cytochrome P450, family 4, subfamily a, polypeptide 14	<i>Cyp4a14</i>	CCCCTCTAGATTTGCACCAGAT TCCAATGCAGTTCCTTGATC	
Acetyl -CoA carboxylase 2	<i>Acacb</i>	GGGCTCCCTGGATGACAAC GCTCTTCCGGGAGGAGTTCT	[7]
Carnitine palmitoyl transferase 1, liver isoform	<i>Cpt1a</i>	GACAGCTATGCCAAATCTCTGCTG CAGGAATGCTCTGCGTTTATGCC	
Carnitine palmitoyl transferase 2	<i>Cpt2</i>	AGCCTACCTGGTCAATGCATATC GGGTTGGGTATACGAGTTGAATT	
Medium-chain specific acyl-CoA dehydrogenase (Mcad)	<i>Acadm</i>	GCAACTGCCCGCAAGTTT TACTCCCCGCTTTTGTTCATATTC	
Long-chain specific acyl-CoA dehydrogenase (Lcad)	<i>Acadl</i>	TCAATGGAAGCAAGGTGTTC GCCACGACGATCACGAGAT	
Microsomal triglyceride transfer protein	<i>Mttp</i>	CCTACCAGGCCCAACAAGAC CGCTCAATTTTGCATGTATCC	[4]
Adipophilin	<i>Adfp</i>	ATGAGTCCCCTGTGTTGAG GCCTGATCTTGAATGTTCTGTG	
Phosphoenolpyruvate carboxykinase, cytosolic	<i>Pck1</i>	GTGCTGGAGTGGATGTTCGG CTGGCTGATTCTCTGTTTCAGG	[8]
Glucose 6 phosphatase	<i>G6pc</i>	ACTGTGGGCATCAATCTCCTC CGGGACAGACAGACGTTTCAGC	[8]
Peroxisome proliferative activated receptor, gamma, coactivator 1 (PGC-1 $\alpha$ )	<i>Ppargc1a</i>	AACCACACCCACAGGATCAGA TCTTCGCTTTATTGCTCCATGA	[4]
Pyruvate dehydrogenase kinase	<i>Pdk1</i>	GTGTTTGCTGAAGCTCCTAAAGG TGTTCAAAACCACGCCCAAT	

## Supplementary References

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