

Supplementary Table S1. Clinical characteristics of the exploratory and validation cohorts.

	Exploratory cohort	Validation cohort	p	Pooled cohort
N	75	39		114
Females (%)	58(77)	33(85)	0.127	91(80)
Age	50.42 ± 10.10	51.82 ± 11.39	0.509	50.53 ± 10.65
BMI	43.43 (41.02, 47.91)	45.97 (40.35, 48.41)	0.565	43.49 (41.01, 48.18)
T2D (%)	21(28)	17(44)	0.155	38(33)
Type of surgery				
SG (%)	27(36)	15(39)	>0.999	42(37)
RYGB (%)	45(60)	24(61)		69(60)
TBWL (%)	31.98 ± 7.97	29.15 ± 6.35	0.043	31.25 ± 7.72

Data are presented as the mean ± SD, median (95%CI) or number (%). BMI, body mass index; T2D, type 2 diabetes; SG, sleeve gastrectomy; RYGB, Roux-en-Y Gastric Bypass; TBWL, total body weight loss.

Supplementary Table S2. Clinical characteristics of non-obese and ex-obese subjects.

	non-Ob (n=17)	ex-Ob (n=18)
Sex (Female)	13 (62%)	16 (89%)
Age	60.81 ± 9.35	49.94 ± 11.87
Weight	73.87 ± 12.16	71.13 ± 7.84
Height	1.65 ± 0.11	1.64 ± 2.12
BMI	27.12 ± 2.55	26.37 ± 2.12
FPG	93 (84, 98)	81.19 ± 8.68
Hypertension	4 (19%)	0 (0%)
Total cholesterol	196.6 ± 35.88	169.0 ± 24.55
HDL	60.05 (55, 84)	55.88 ± 11.57
LDL	104.5 ± 33.53	97.25 ± 17.94
TG	98.23 ± 37.93	79.19 ± 22.07
AST	20.56 ± 6.74	20 (17, 29)
ALT	18 (14, 36)	21 (10, 38)
GGT	32.3 ± 14.21	10.5 (8, 27)

Data are presented as the mean \pm SD, median (95%CI) or number (%). BMI, body mass index; FPG, fasting plasma glucose; HDL, serum high-density lipoprotein cholesterol; LDL, serum low-density lipoprotein cholesterol; TG, serum triglycerides; AST, serum aspartate aminotransferase; ALT, serum alanine aminotransferase; GGT, gamma-glutamyl transferase.

Supplementary Table S3. List of primers used in the study.

GENE	SEQUENCE	(5'-3')	GENE	SEQUENCE	(5'-3')
ABCA1	F	GGAGGCCAGAACATGACATCTTAG	LOX	F	ACATTGCGTACACAGGACATC
ABCA1	R	TTTCAGCCCCATTAACTCC	LOX	R	TCCCCACTTCAGAACACCAG
PLIN2	F	AGTATCCCTACCTGAAAGTCTGTG	LOXL2	F	GTGCAGCGACAAAAGGATTG
PLIN2	R	CCCCCTACAGGCATAGGTATTG	LOXL2	R	GCGGTAGGTGAGAGGTATG
ADIPOQ	F	ACAATGACTCCACCTCACAG	LOXL4	F	GTGGCAGAGTCAGATTTCTCC
ADIPOQ	R	TTCTAACCGTACTGAAGGCC	LOXL4	R	TTGTTCTGAGACGCTGTTC
ADIPOR1	F	TCACTACCTCCATCGCTGT	LPL	F	GGACTGAGAGTGAAACCCATAC
ADIPOR1	R	CACTAAGCCAAGTCCCAG	LPL	R	GGAAGGAGTAGGTCTTATTTGTGG
ADIPOR2	F	CAGCCATTATACTGCTCCCAGTG	MCP1 / CCL2	F	CCTCAGCATGAAAGTCTCTG
ADIPOR2	R	CCGAGATGACATAGTGAAGG	MCP1 / CCL2	R	TCTGCACTGAGATCTCCATTG
ANGPT1	F	TTAAAGGACTTACAGGGACAGC	MMP13	F	GGAATTGGTATAAAAGTAGATGCTG
ANGPT1	R	GACACATGCAACCAACAC	MMP13	R	ACGGAAACAATACGGTTACTC
ANGPT2	F	CCAGGAGACTTGAACCTCAGC	MMP14	F	GGATGGACACGGAGAAATTG
ANGPT2	R	TGTGTTGCTTCATAGCTAG	MMP14	R	TTTACAGGAACAGAAAGGCCG
ATG12	F	AATAGTCTTGTCTCTTCC	MMP15	F	CCAAGCCATCAGTGTGTC
ATG12	R	GCAAGTGTATTTCTGGTGTTC	MMP15	R	CGCTTATTGCGAATTTCAG
ATG5	F	AGCAACTCTGAGTGGATTG	MMP2	F	ACCCATTACACCTACACCAAAG
ATG5	R	AGGTCTTCAGTCGGTGTCTG	MMP2	R	TGTTGAGATCTCAGGAGTG
ATG7	F	TTTGCTATCTGCCCTCTG	MMP9	F	CGAACATTGACAGCAGAACAG
ATG7	R	GCTGTAACCTCTTGTGAC	MMP9	R	CACTGAGGAATGATCTAAGGCC
ATGL	F	CACTCAACTCCAAGGACGAG	MOGAT1	F	GAAAGCCATCCACACTGTG
ATGL	R	CTCATAGTGGCAGGTGTGTC	MOGAT1	R	GCCATACTTCTTGTGTTCC
BGN	F	TGGAAACAGTGGCTTGAAAC	MSR1	F	ATCTGTAATTTGATGCTCGC
BGN	R	GTTGTTGCTAGGTGGAGTT	MSR1	R	CCAATGAGAGGATGAGAACTG
CD14	F	CAGAGGCTGGAAAGACTATCG	NPY1R	F	GATCTGAGGAGGAAATACAG
CD14	R	TTGGAGAAGTGTGAGACG	NPY1R	R	CTGAATTCTCATTCCTGAACTG
CD206 / MRC1	F	GCAAAGTGGATTCTGTCCTG	p16 / CDKN2A	F	GATGTGCAAGGGTACCTG
CD206 / MRC1	R	CTGTTATGCTGGCTGCAAATG	p16 / CDKN2A	R	TCTCTGGTTCTTCATCGGG
CD68	F	ATGGGGTGGAGTACAATG	P21 / CDKN1A	F	GAACCTCGACTTTGTCACCGGAGC
CD68	R	TGGACAGCTGGTAAAGAATG	P21 / CDKN1A	R	TGGAGTGGTAGAAATCTGTCATGCT
COL1A1	F	CCCCCTGGAAAATGGAGATG	p53 / TP53	F	CAGCACATGACGGAGGTGT
COL1A1	R	CATCCAACCACTGAAACCTC	p53 / TP53	R	TCATCCAAATACTCCACACGC
COL3A1	F	ATGGTGCTCTGGTAAGAATG	PAI-1 / SERPINE1	F	GTGGACTTTGAGGGTGGAG
COL3A1	R	GGTCTCTGTCCTCTTGT	PAI-1 / SERPINE1	R	GAAGTAGAGGGCATTCCACAG
COL4A1	F	TGTTGATGGCTACTCTTG	PDGFRA	F	TTCTCTGCTGACATTGAC
COL4A1	R	TAGTAAATTGAGGTCCACAG	PDGFRA	R	GTCCTCAATTGGCTGTCCTC
COL5A1	F	TCGCTTACAGAGTCACCAAAG	PDGFRB	F	ATGTGACGGAGAGTGTGAATG
COL5A1	R	GTTTAGATGGAGACCGAGGAAG	PDGFRB	R	GCAGCTCAGCAAATTGTAGTG
COL6A1	F	AGGAGTCAGGAGGAAAG	PLIN1	F	CATTGAGAAAGGTGGTGGAGTAC
COL6A1	R	GCATTACAGCAAGGAC	PLIN1	R	GTGTATCGAGAGAGGGTTG
COL6A3	F	AAACATTGGCTCTACTGAAACA	PLIN2	F	AGTATCCCTACCTGAAGTCTG
COL6A3	R	GCACAGCTTGGTGTGTTGA	PLIN2	R	CCCCTTACAGGCATAGGTATTG
DGAT2	F	TCCGAAATGCTGTGTTGAG	PPARG	F	GTCGGTTTCAAGGATCTG
DGAT2	R	CAAATAGTCTATGGTGGCCCG	PPARG	R	GCTGGTCGATATACTGGAG
ELN	F	CCTGGCTTCGGATTGTC	PPARGC1A / PGC1A	F	CAGGCACTGAGTCTTCAG
ELN	R	CAAAGGGTTACATTCCACC	PPARGC1A / PGC1A	R	TCCTCGTAGTGTACACCTG
F13A1	F	ATCCCATCAAGTCAGCG	RPL6 F	F	CTTAAATTCTTCTTCCATTTG
F13A1	R	ATGTCAAGCTTCAGTC	RPL6 R	R	TTCTGGCTTGGGTTCTT
FABP4	F	CATGTGAGAAATGGGATGG	SLC2A1 / GLUT1	F	TCTATGCTGCTGAACTCTTC
FABP4	R	AACTTCAGTCAGTCACG	SLC2A1 / GLUT1	R	GATGAAGACGTAGGGACAC
FASN	F	CAGAGTCGGAGAACCTGCG	SLC2A4 / GLUT4	F	ACTGGACGAGCAACTTCATC
FASN	R	GGAGGCATCAACCTAGACAG	SLC2A4 / GLUT4	R	GAGGACCCGAAATAGAAGGAA
FN1	F	GTGGAGAAAGAATATCTCG	SREBF1	F	TTCTGACAGCCATGAAGACAG
FN1	R	GAGAATACTGGTTAGGACTGG	SREBF1	R	CCGCATCTACGACAGT
HAS1	F	CTTGTAGAGCTACTCCACTG	TCF21	F	ATACGAGAACGGTACATTAC
HAS1	R	CGGTATCCCCAAAAGTACAG	TCF21	R	CTCGCTCCAGGTACCAAAC
HAS2	F	CTCAATTGGAAATGCGCCG	TGFB1	F	TTGATGTACCGGGAGTTG
HAS2	R	CTCAGGATACATAAGAACCTCTAC	TGFB1	R	GTAGTAAACCCGTTGATGTC
HIF1A	F	AAAGAACTTAGGGCGCTCA	TIMP1	F	CCCCTAGCGTGGACATTATC
HIF1A	R	CAACCCAGACATATCCACCTC	TIMP1	R	AAACAGGATGCCAGAAGCC
HSL / LIPE	F	TCATCTCCATGACTACTCC	TIMP2	F	CCCCCTGTGACTTCATCG
HSL / LIPE	R	AGATTGCTTCCCCTGTGAG	TIMP2	R	GAGATGTAGCACGGGATCATG
HYAL1	F	GATGGCTGTGGAGTCAATG	TIMP3	F	CTCTGCAACTCCGACATC
HYAL1	R	CCCAAGACTAGTTCT	TIMP3	R	GCCTCGGTACATCTCATCG
HYAL2	F	TGTGAGCTTCGGTGT	TNFA	F	AGGTCTACTTGGGATCTGC
HYAL2	R	CTCGCAATGGTGGAGATGAG	TNFA	R	GAAGAGGTTGAGGGTCTG
IL1B	F	GGTACATCAGCACCTCAAG	UCP1	F	GGACTACTCCAAATCTGATGAG
IL1B	R	CACATTCAAGCAGGACTCTC	UCP1	R	AAATCCAGGATAAGAGCCG
IL6	F	CAACCTGAACCTTCAAAAGATG	UCP2	F	TCCTGAAAGCCAACCTCATG
IL6	R	ACCTCAACTTCAAAAGACCGAG	UCP2	R	GGCAGAGTTCATGATCTGTC
IRS1	F	TCTGCTCAGGGTGGGT	UCP3	F	AGAAAATACAGCGGGACTATGG
IRS1	R	GTGCATGCTTGGGTT	UCP3	R	CTTGAGGATGTCGTAGGTAC
ITGB2	F	CCCCAAGTTGCTGAGAGTTAG	VEGFA	F	AGTCCAACATCACCAGCAG
ITGB2	R	TGAGGGCGGAAATACTGG	VEGFA	R	TTCCCTTCCCTGAACTGATT
KLB	F	CATGGGTATGGGACAGGTATG	VEGFB	F	CTTAGAGCTCAACCCAGACAC
KLB	R	TCTGATGTGGCGGAAATG	VEGFB	R	ACCCCTGCTGAGTGTGAAAAG
LEP	F	GCTTCAGGCTACTCCACAG	VEGFR1	F	TCCCTCAACCTACAACTAAGTG
LEP	R	CCTTCCCTTAACGTAGTCCITG	VEGFR1	R	GCTCTCAATTCTGTTCCATG
LEPR	F	TCAACAGTACAATCCAGTCAC	VEGFR2	F	CATTCAAAAGGAGAAGCAGAC
LEPR	R	TTTGGGCTAGATATGGGATG	VEGFR2	R	GAGGAATGGCATAGACCGTAC

F, forward; R, reverse.

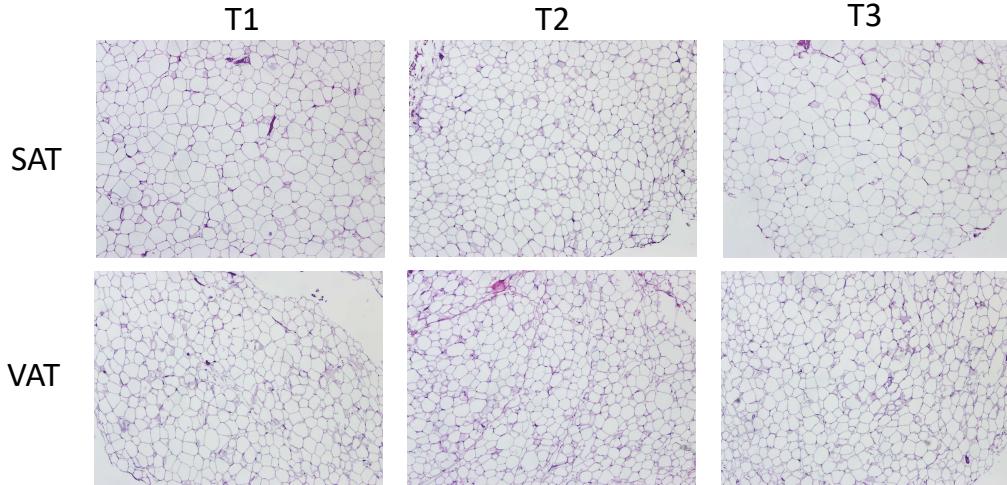
Supplementary Table S4. Summary of multivariate regression models predicting TBWL.

Model	Predictor	β	SE	95% CI	p	Adjusted R ²	p
1	Constant	29.394	1.539	(26.35, 32.44)	<0.001	0.134	<0.001
	Type of BS	4.387	1.365	(1.69, 7.09)	0.002		
	Age	-0.128	0.061	(-0.25, -0.01)	0.038		
	Sex	-0.045	1.369	(-2.75, 2.66)	0.974		
	BMI	0.079	0.125	(-0.17, 0.33)	0.529		
2	T2D	-2.543	1.347	(-5.21, 0.12)	0.061	0.228	<0.001
	Constant	25.888	2.085	(21.75, 30.03)	<0.001		
	Type of BS	6.689	1.749	(3.21, 10.17)	<0.001		
	Age	-0.192	0.075	(-0.34, -0.04)	0.012		
	Sex	1.595	1.619	(-1.62, 4.81)	0.327		
	BMI	0.06	0.153	(-0.25, 0.37)	0.697		
3	T2D	-1.614	1.567	(-4.73, 1.5)	0.306	0.306	<0.001
	COL1A1	-5.501	1.909	(-9.3, -1.71)	0.005		
	Constant	26.41	2.048	(22.33, 30.49)	<0.001		
	Type of BS	5.424	1.827	(1.79, 9.06)	0.004		
	Age	-0.146	0.076	(-0.3, 0.01)	0.059		
4	Sex	2.293	1.593	(-0.88, 5.47)	0.154	0.301	<0.001
	BMI	0.145	0.155	(-0.16, 0.45)	0.353		
	T2D	-1.659	1.542	(-4.73, 1.41)	0.286		
	COL5A1	-24.732	5.58	(-35.84, -13.62)	<0.001		
	Constant	27.128	1.791	(23.57, 30.69)	<0.001		
5	Type of BS	6.015	1.603	(2.83, 9.2)	<0.001	0.301	<0.001
	Age	-0.167	0.069	(-0.3, -0.03)	0.018		
	Sex	1.434	1.48	(-1.51, 4.38)	0.335		
	BMI	0.095	0.142	(-0.19, 0.38)	0.503		
	T2D	-2.301	1.437	(-5.16, 0.56)	0.113		
6	COL6A3	-12.712	2.741	(-18.16, -7.26)	<0.001	0.301	<0.001
	Constant	27.128	1.791	(23.57, 30.69)	<0.001		

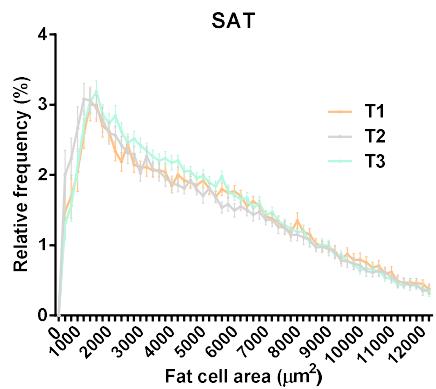
Type of BS: 0-SG, 1-GBP; sex: 0-male: 1-female; T2D: 0-no; 1-yes. β : Beta coefficient; SE: standard error; CI: confidence interval.

Supplementary Figure S1. Adipocyte size distribution across weight loss tertiles.

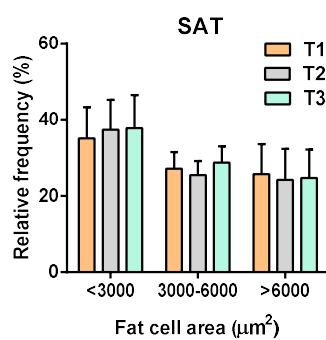
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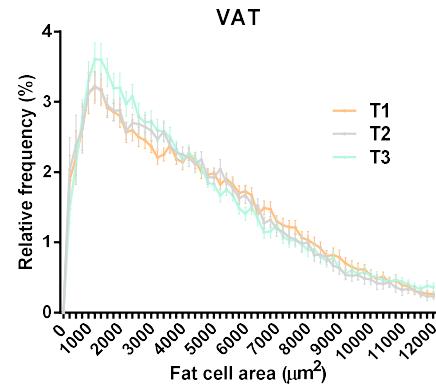
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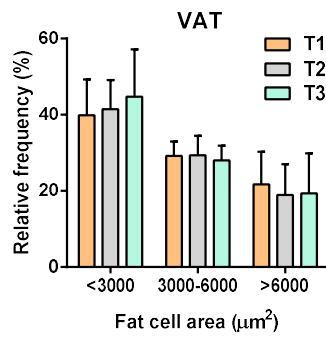
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D



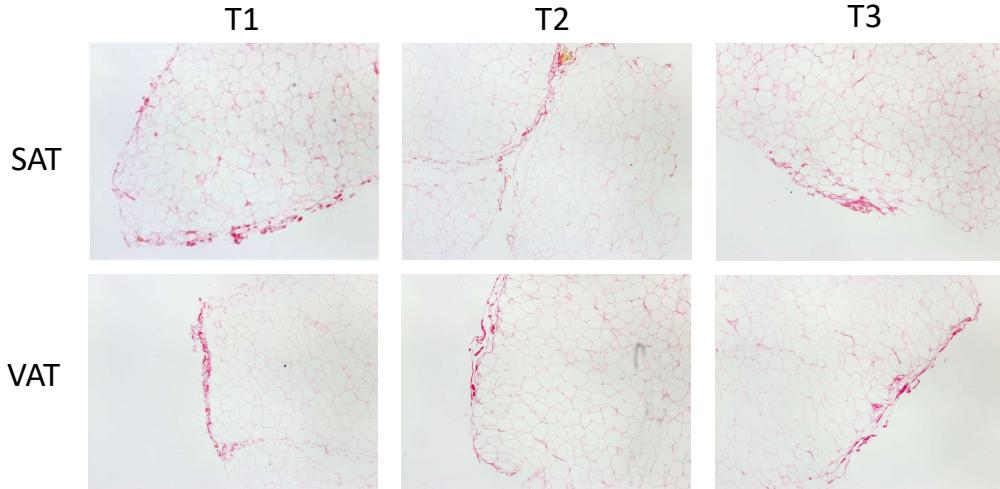
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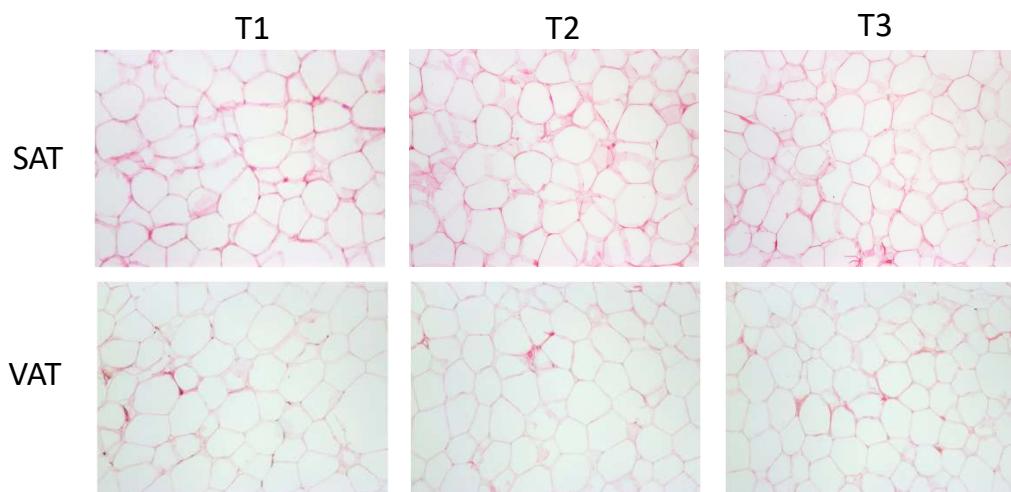
A: Representative images of hematoxylin and eosin staining of SAT and VAT samples among TBWL tertile groups. B, C: Frequency distribution analysis of fat cell areas divided by size into bin intervals of $200 \mu\text{m}^2$ (B) and three representative sizes in SAT (C). D, E: Frequency distribution analysis of fat cell areas divided by size into bin intervals of $200 \mu\text{m}^2$ (D) and three representative sizes in VAT (E). Data are presented as the mean \pm SD.

Supplementary Figure S2. Adipose tissue fibrosis across weight loss tertiles.

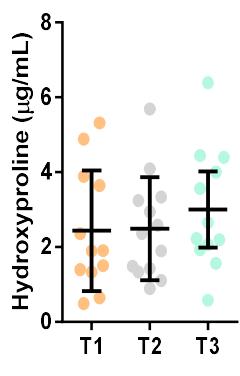
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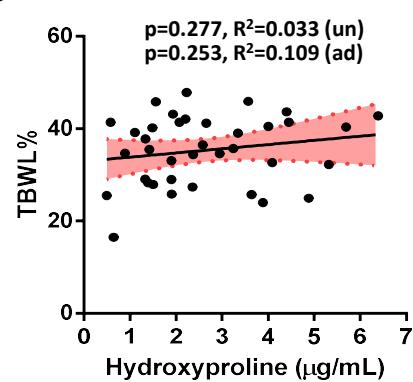
B



C



D



A, B: Representative images of Sirius red staining showing total (A) and pericellular fibrosis (B) in SAT and VAT samples among TBWL tertile groups. C: Determination of SAT hydroxyproline residues in TBWL tertiles. Data are presented as the mean \pm SD. D: Regression analysis between hydroxyproline concentration in SAT and TBWL%. Simple linear regression (solid line) and 95% confidence interval (dashed lines) are shown. P-values are unadjusted (un) or adjusted for sex, age, baseline BMI, T2D and type of surgery (ad).