

Table S1. Gene ontology using up-regulated genes in sarcopenia (cutoff: P value <0.05)

Category	Term	Count	%	P value	Genes
GOTERM_BP_FAT	GO:0002011~morphogenesis of an epithelial sheet	5	2.702703	0.001	VANGL2, FLRT3, AJUBA, BMP7, CYR61
GOTERM_BP_FAT	GO:0030198~extracellular matrix organization	11	5.945946	0.001	HPSE2, ELF3, ITGA3, ITGA2, COL4A6, COL4A5, COL9A3, LAMC2, ITGB6, THSD4, CYR61
GOTERM_BP_FAT	GO:0043062~extracellular structure organization	11	5.945946	0.002	HPSE2, ELF3, ITGA3, ITGA2, COL4A6, COL4A5, COL9A3, LAMC2, ITGB6, THSD4, CYR61
GOTERM_BP_FAT	GO:0007167~enzyme linked receptor protein signaling pathway	20	10.81081	0.002	KL, IRS1, CAV2, ITGA3, HFE, ARHGEF28, NRG1, BMP7, PRLR, DKK1, CYR61, DNMT1, FLRT3, CHN1, COL4A5, SPRY1, ROR1, ATP6V0A4, RNF165, ATP6V1C2
GOTERM_BP_FAT	GO:0007409~axonogenesis	12	6.486486	0.002	LRFN5, VANGL2, BCL11A, FLRT3, CHN1, NRXN3, SLTRK6, NTN4, NRG1, BMP7, RNF165, ROBO1
GOTERM_BP_FAT	GO:0006805~xenobiotic metabolic process	6	3.243243	0.003	GSTM2, GSTM1, FMO2, CYP3A5, UGT1A6, CES1
GOTERM_BP_FAT	GO:0048667~cell morphogenesis involved in neuron differentiation	13	7.027027	0.003	BCL11A, NTN4, NRXN3, NRG1, BMP7, ROBO1, LRFN5, VANGL2, FLRT3, ID2, CHN1, SLTRK6, RNF165
GOTERM_BP_FAT	GO:0070886~positive regulation of calcineurin-NFAT signaling cascade	3	1.621622	0.003	AKAP6, NRG1, LMCD1
GOTERM_BP_FAT	GO:0071466~cellular response to xenobiotic stimulus	6	3.243243	0.003	GSTM2, GSTM1, FMO2, CYP3A5, UGT1A6, CES1
GOTERM_BP_FAT	GO:046578~regulation of Ras protein signal transduction	8	4.324324	0.004	RASA4B, ITGA3, RASA4, ARHGEF28, ARHGEF4, SPRY1, NRG1, ROBO1
GOTERM_BP_FAT	GO:0007265~Ras protein signal transduction	10	5.405405	0.004	VANGL2, RASA4B, ITGA3, RASA4, PLK2, ARHGEF28, ARHGEF4, SPRY1, NRG1, ROBO1
GOTERM_BP_FAT	GO:0061564~axon development	12	6.486486	0.004	LRFN5, VANGL2, BCL11A, FLRT3, CHN1, NRXN3, SLTRK6, NTN4, NRG1, BMP7, RNF165, ROBO1
GOTERM_BP_FAT	GO:0009410~response to xenobiotic stimulus	6	3.243243	0.004	GSTM2, GSTM1, FMO2, CYP3A5, UGT1A6, CES1
GOTERM_BP_FAT	GO:0017144~drug metabolic process	4	2.162162	0.005	CYP4B1, FMO2, CYP4F12, CYP3A5
GOTERM_BP_FAT	GO:0007264~small GTPase mediated signal transduction	13	7.027027	0.005	ARL10, RASA4B, ITGA3, PLK2, ARHGEF28, NRG1, RND3, ROBO1, VANGL2, RASA4, ARHGEF4, RAB38, SPRY1
GOTERM_BP_FAT	GO:0031581~hemidesmosome assembly	3	1.621622	0.006	DST, LAMC2, KRT5
GOTERM_BP_FAT	GO:0007411~axon guidance	8	4.324324	0.006	VANGL2, FLRT3, CHN1, NRXN3, NTN4, BMP7, RNF165, ROBO1
GOTERM_BP_FAT	GO:0097485~neuron projection guidance	8	4.324324	0.006	VANGL2, FLRT3, CHN1, NRXN3, NTN4, BMP7, RNF165, ROBO1
GOTERM_BP_FAT	GO:0007369~gastrulation	7	3.783784	0.006	OSR2, VANGL2, DUSP1, ITGA3, ITGA2, BMP7, DKK1
GOTERM_BP_FAT	GO:0060429~epithelium development	20	10.81081	0.006	KRT4, ITGA2, NTN4, NRG1, BMP7, PRLR, DKK1, CYR61, ROBO1, VANGL2, ELF3, FLRT3, KRT16, ID2, SLTRK6, SPRY1, ROR1, AJUBA, CES1, KRT6A
GOTERM_BP_FAT	GO:0050850~positive regulation of calcium-mediated signaling	4	2.162162	0.007	GSTM2, AKAP6, NRG1, LMCD1
GOTERM_BP_FAT	GO:0050808~synapse organization	8	4.324324	0.007	NLG4Y, CACNB2, FLRT3, NRXN3, SLTRK6, COL4A5, NRG1, DKK1
GOTERM_BP_FAT	GO:0048729~tissue morphogenesis	14	7.567568	0.007	ITGA3, ITGA2, NTN4, NRG1, BMP7, DKK1, CYR61, VANGL2, FLRT3, KRT16, SPRY1, ROR1, AJUBA, KRT6A
GOTERM_BP_FAT	GO:0046580~negative regulation of Ras protein signal transduction	4	2.162162	0.007	RASA4B, ITGA3, RASA4, SPRY1
GOTERM_BP_FAT	GO:0042330~taxis	13	7.027027	0.007	ITGA2, NTN4, NRXN3, NRG1, BMP7, CYR61, ROBO1, VANGL2, FLRT3, ID2, CHN1, S100A14, RNF165
GOTERM_BP_FAT	GO:0051058~negative regulation of small GTPase mediated signal transduction	4	2.162162	0.008	RASA4B, ITGA3, RASA4, SPRY1
GOTERM_BP_FAT	GO:0051056~regulation of small GTPase mediated signal transduction	9	4.864865	0.008	RASA4B, ITGA3, RASA4, ARHGEF28, CHN1, ARHGEF4, SPRY1, NRG1, ROBO1
GOTERM_BP_FAT	GO:0008544~epidermis development	9	4.864865	0.009	VANGL2, ELF3, KRT16, SLTRK6, LAMC2, KRT5, DKK1, EDA2R, ZNF750
GOTERM_BP_FAT	GO:0030879~mammary gland development	6	3.243243	0.009	ELF3, ID2, ITGA2, NRG1, PRLR, ROBO1
GOTERM_BP_FAT	GO:0048598~embryonic morphogenesis	13	7.027027	0.009	OSR2, DUSP1, ITGA3, ITGA2, FOXI1, BMP7, DKK1, CYR61, VANGL2, FLRT3, ID2, SLTRK6, SP8
GOTERM_BP_FAT	GO:0007169~transmembrane receptor protein tyrosine kinase signaling pathway	14	7.567568	0.010	KL, IRS1, CAV2, ARHGEF28, NRG1, PRLR, DNM1, FLRT3, CHN1, COL4A5, SPRY1, ROR1, ATP6V0A4, ATP6V1C2
GOTERM_BP_FAT	GO:0071495~cellular response to endogenous stimulus	21	11.35135	0.010	KL, GSTM2, DUSP1, IRS1, CAV2, ITGA3, HFE, LMO3, AKAP6, ROR8, BMP7, PRLR, DKK1, CYR61, CPEB1, FLRT3, COL4A6, SPRY1, ATP6V0A4, RNF165, ATP6V1C2
GOTERM_BP_FAT	GO:0007044~cell-substrate junction assembly	5	2.702703	0.010	DST, ITGA2, LAMC2, KRT5, AJUBA
GOTERM_BP_FAT	GO:0009790~embryo development	18	9.72973	0.010	OSR2, DUSP1, ITGA3, NRK, ITGA2, FOXI1, NRG1, BMP7, DKK1, CYR61, STOX2, KRT19, VANGL2, ELF3, FLRT3, ID2, SLTRK6, SP8
GOTERM_BP_FAT	GO:0070887~cellular response to chemical stimulus	38	20.54054	0.011	IRS1, HFE, AKAP6, ROR8, CYP3A5, CYR61, EDA2R, ROBO1, DUOX1, FLRT3, ATP6V0A4, S100A14, PRODH, ATP6V1C2, UGT1A6, KL, GSTM2, GSTM1, CAV2, ITGA3, DUSP1, LMO3, ITGA2, KRT13, TNFRSF10C, FMO2, BMP7, DKK1, PRLR, IL17RB, CPEB1, CYP24A1, RASA4, ID2, COL4A6, SPRY1, RNF165, CES1
GOTERM_BP_FAT	GO:0070884~regulation of calcineurin-NFAT signaling cascade	3	1.621622	0.011	AKAP6, NRG1, LMCD1
GOTERM_BP_FAT	GO:0051962~positive regulation of nervous system development	11	5.945946	0.012	BCL11A, FLRT3, ITGA3, ID2, DUOX1A1, NRXN3, SLTRK6, NRG1, BMP7, DKK1, ROBO1
GOTERM_BP_FAT	GO:1902532~negative regulation of intracellular signal transduction	11	5.945946	0.012	GSTM2, RASA4B, DUSP1, FLRT3, ITGA3, RASA4, LMO3, WWC2, SPRY1, AJUBA, BMP7
GOTERM_BP_FAT	GO:1902531~regulation of intracellular signal transduction	27	14.59459	0.013	IRS1, ARHGEF28, NRK, WWC2, AKAP6, CYR61, EDA2R, ROBO1, FLRT3, CHN1, LMCD1, KL, GSTM2, RASA4B, CAV2, ITGA3, DUSP1, LMO3, PLK2, NRG1, BMP7, PRLR, VANGL2, RASA4, ARHGEF4, SPRY1, AJUBA
GOTERM_BP_FAT	GO:0042537~benzene-containing compound metabolic process	3	1.621622	0.016	GSTM2, GSTM1, CYP4B1
GOTERM_BP_FAT	GO:0048812~neuron projection morphogenesis	12	6.486486	0.016	LRFN5, VANGL2, BCL11A, FLRT3, CHN1, NRXN3, SLTRK6, NTN4, NRG1, BMP7, RNF165, ROBO1
GOTERM_BP_FAT	GO:0051960~regulation of nervous system development	15	8.108108	0.016	BCL11A, ITGA3, PLK2, DIXDC1, NRXN3, NRG1, BMP7, DKK1, ROBO1, VANGL2, FLRT3, ID2, DUOX1A1, CHN1, SLTRK6
GOTERM_BP_FAT	GO:0090092~regulation of transmembrane receptor protein serine/threonine kinase signaling pathway	7	3.783784	0.017	CAV2, ITGA3, HFE, BMP7, DKK1, CYR61, RNF165
GOTERM_BP_FAT	GO:0033173~calcineurin-NFAT signaling cascade	3	1.621622	0.017	AKAP6, NRG1, LMCD1
GOTERM_BP_FAT	GO:0009968~negative regulation of signal transduction	19	10.27027	0.017	GSTM2, RASA4B, DUSP1, IRS1, CAV2, ITGA3, LMO3, WWC2, DDIT4L, NRG1, KIF7, BMP7, DKK1, DNM1, ROBO1, RASA4, FLRT3, SPRY1, AJUBA
GOTERM_BP_FAT	GO:0006935~chemotaxis	12	6.486486	0.018	VANGL2, FLRT3, ITGA2, CHN1, NRXN3, NTN4, NRG1, S100A14, BMP7, CYR61, RNF165, ROBO1
GOTERM_BP_FAT	GO:0009967~positive regulation of signal transduction	23	12.43243	0.018	KL, GSTM2, IRS1, CAV2, HFE, LMO3, PLK2, NRK, DIXDC1, AKAP6, NRG1, KIF7, BMP7, PRLR, CYR61, EDA2R, ROBO1, VANGL2, CHN1, AJUBA, LMCD1, RNF165, ATP6V1C2
GOTERM_BP_FAT	GO:1901687~glutathione derivative biosynthetic process	3	1.621622	0.019	GSTM2, GSTM1, GSTMS
GOTERM_BP_FAT	GO:1901685~glutathione derivative metabolic process	3	1.621622	0.019	GSTM2, GSTM1, GSTMS
GOTERM_BP_FAT	GO:0010648~negative regulation of cell communication	20	10.81081	0.019	GSTM2, RASA4B, DUSP1, IRS1, CAV2, ITGA3, LMO3, PLK2, WWC2, DDIT4L, NRG1, KIF7, BMP7, DKK1, DNM1, ROBO1, RASA4, FLRT3, SPRY1, AJUBA

GOTERM_BP_FAT	GO:0023057~negative regulation of signaling	20	10.81081	0.019	<i>GSTM2, RASA4B, DUSP1, IRS1, CAV2, ITGA3, LMO3, PLK2, WWC2, DDIT4L, NRG1, KIF7, BMP7, DKK1, DNMT1, ROBO1, RASA4A, FLRT3, SPRY1, AJUBA</i>
GOTERM_BP_FAT	GO:0010243~response to organonitrogen compound	15	8.108108	0.020	<i>KL, GSTM2, DUSP1, IRS1, CAV2, ITGA2, AKAP6, BMP7, PRLR, DUOX1, CPEB1, PPP1R1B, COL4A6, ATP6V0A4, ATP6V1C2</i>
GOTERM_BP_FAT	GO:0060411~cardiac septum morphogenesis	4	2.162162	0.023	<i>VANGL2, ID2, BMP7, CYR61</i>
GOTERM_BP_FAT	GO:0009719~response to endogenous stimulus	24	12.97297	0.023	<i>KL, GSTM2, DUSP1, IRS1, CAV2, ITGA3, HFE, LMO3, ITGA2, AKAP6, RORR, BMP7, PRLR, DKK1, CYR61, DUOX1, CPEB1, FLRT3, PPP1R1B, COL4A6, SPRY1, ATP6V0A4, RNF165, ATP6V1C2</i>
GOTERM_BP_FAT	GO:0008286~insulin receptor signaling pathway	5	2.702703	0.024	<i>KL, IRS1, CAV2, ATP6V0A4, ATP6V1C2</i>
GOTERM_BP_FAT	GO:0010647~positive regulation of cell communication	24	12.97297	0.025	<i>KL, GSTM2, IRS1, CAV2, HFE, LMO3, PLK2, NRK, ITGA2, DIXDC1, AKAP6, NRG1, KIF7, BMP7, PRLR, CYR61, EDA2R, ROBO1, VANGL2, CHN1, AJUBA, LMCD1, RNF165, ATP6V1C2</i>
GOTERM_BP_FAT	GO:1901698~response to nitrogen compound	16	8.648649	0.026	<i>KL, GSTM2, GSTM1, DUSP1, IRS1, CAV2, ITGA2, AKAP6, BMP7, PRLR, DUOX1, CPEB1, PPP1R1B, COL4A6, ATP6V0A4, ATP6V1C2</i>
GOTERM_BP_FAT	GO:2000725~regulation of cardiac muscle cell differentiation	3	1.621622	0.026	<i>AKAP6, NRG1, DKK1</i>
GOTERM_BP_FAT	GO:0023056~positive regulation of signaling	24	12.97297	0.026	<i>KL, GSTM2, IRS1, CAV2, HFE, LMO3, PLK2, NRK, ITGA2, DIXDC1, AKAP6, NRG1, KIF7, BMP7, PRLR, CYR61, EDA2R, ROBO1, VANGL2, CHN1, AJUBA, LMCD1, RNF165, ATP6V1C2</i>
GOTERM_BP_FAT	GO:0007507~heart development	11	5.945946	0.026	<i>FHOD3, VANGL2, FLRT3, ITGA3, ID2, AKAP6, NRG1, BMP7, DKK1, CYR61, ROBO1</i>
GOTERM_BP_FAT	GO:0001704~formation of primary germ layer	5	2.702703	0.027	<i>DUSP1, ITGA3, ITGA2, BMP7, DKK1</i>
GOTERM_BP_FAT	GO:0003206~cardiac chamber morphogenesis	5	2.702703	0.027	<i>VANGL2, ID2, NRG1, BMP7, CYR61</i>
GOTERM_BP_FAT	GO:0045995~regulation of embryonic development	5	2.702703	0.027	<i>OSR2, VANGL2, NRK, BMP7, DKK1</i>
GOTERM_BP_FAT	GO:0090383~phagosome acidification	3	1.621622	0.028	<i>RAB38, ATP6V0A4, ATP6V1C2</i>
GOTERM_BP_FAT	GO:0070458~cellular detoxification of nitrogen compound	2	1.081081	0.029	<i>GSTM2, GSTM1</i>
GOTERM_BP_FAT	GO:0001707~mesoderm formation	4	2.162162	0.029	<i>ITGA3, ITGA2, BMP7, DKK1</i>
GOTERM_BP_FAT	GO:0000904~cell morphogenesis involved in differentiation	14	7.567568	0.030	<i>BCL11A, NTN4, NRXN3, NRG1, BMP7, DKK1, ROBO1, LRFN5, VANGL2, FLRT3, ID2, CHN1, SLTRK6, RNF165</i>
GOTERM_BP_FAT	GO:0048332~mesoderm morphogenesis	4	2.162162	0.031	<i>ITGA3, ITGA2, BMP7, DKK1</i>
GOTERM_BP_FAT	GO:0030278~regulation of ossification	6	3.243243	0.031	<i>KL, OSR2, ID2, RORR, BMP7, CYR61</i>
GOTERM_BP_FAT	GO:0048738~cardiac muscle tissue development	6	3.243243	0.034	<i>FHOD3, ID2, AKAP6, NRG1, BMP7, DKK1</i>
GOTERM_BP_FAT	GO:0090596~sensory organ morphogenesis	7	3.783784	0.034	<i>OSR2, VANGL2, KRT13, FOXI1, SLTRK6, RORR, BMP7</i>
GOTERM_BP_FAT	GO:0032869~cellular response to insulin stimulus	6	3.243243	0.035	<i>KL, CPEB1, IRS1, CAV2, ATP6V0A4, ATP6V1C2</i>
GOTERM_BP_FAT	GO:1901021~positive regulation of calcium ion transmembrane transporter activity	3	1.621622	0.036	<i>CACNB2, GSTM2, AKAP6</i>
GOTERM_BP_FAT	GO:0050890~cognition	7	3.783784	0.036	<i>NLGN4Y, ITGA3, PLK2, PPP1R1B, NRXN3, TMPRSS11E, DKK1</i>
GOTERM_BP_FAT	GO:0007416~synapse assembly	5	2.702703	0.037	<i>NLGN4Y, FLRT3, NRXN3, SLTRK6, NRG1</i>
GOTERM_BP_FAT	GO:0044708~single-organism behavior	9	4.864865	0.037	<i>NLGN4Y, ITGA3, PLK2, ID2, PPP1R1B, NRXN3, SLTRK6, DKK1, DNMT1</i>
GOTERM_BP_FAT	GO:0048333~mesodermal cell differentiation	3	1.621622	0.038	<i>ITGA3, ITGA2, DKK1</i>
GOTERM_BP_FAT	GO:0048016~inositol phosphate-mediated signaling	3	1.621622	0.038	<i>AKAP6, NRG1, LMCD1</i>
GOTERM_BP_FAT	GO:0018916~nitrobenzene metabolic process	2	1.081081	0.038	<i>GSTM2, GSTM1</i>
GOTERM_BP_FAT	GO:1900138~negative regulation of phospholipase A2 activity	2	1.081081	0.038	<i>ANXA8L1, ANXA8</i>
GOTERM_BP_FAT	GO:0050848~regulation of calcium-mediated signaling	4	2.162162	0.039	<i>GSTM2, AKAP6, NRG1, LMCD1</i>
GOTERM_BP_FAT	GO:0050877~neurological system process	20	10.81081	0.040	<i>NLGN4Y, PRRT2, PDE1C, ITGA3, PLK2, ITGA2, NRXN3, OR5P3, NRG1, OR5P2, RORB, TMPRSS11E, DKK1, DNMT1, SCNN1G, CACNB2, PPP1R1B, SLTRK6, ATP6V0A4, TRPM3</i>
GOTERM_BP_FAT	GO:0071417~cellular response to organonitrogen compound	10	5.405405	0.041	<i>KL, GSTM2, CPEB1, IRS1, CAV2, COL4A6, AKAP6, ATP6V0A4, PRLR, ATP6V1C2</i>
GOTERM_BP_FAT	GO:0003156~regulation of organ formation	3	1.621622	0.042	<i>SPRY1, BMP7, DKK1</i>
GOTERM_BP_FAT	GO:0044255~cellular lipid metabolic process	17	9.189189	0.043	<i>KL, GSTM2, ST6GALNAC2, INPP1, IRS1, ACOT11, NRG1, CYP3A5, CYP4F12, CYR61, ACSF2, ST3GAL5, SPTSSB, BC02, AJUBA, CES1, UGT1A6</i>
GOTERM_BP_FAT	GO:0030509~BMP signaling pathway	5	2.702703	0.043	<i>HFE, BMP7, DKK1, CYR61, RNF165</i>
GOTERM_BP_FAT	GO:0045664~regulation of neuron differentiation	11	5.945946	0.044	<i>BCL11A, ITGA3, PLK2, ID2, DUOX1, CHN1, DIXDC1, NRG1, BMP7, DKK1, ROBO1</i>
GOTERM_BP_FAT	GO:0030510~regulation of BMP signaling pathway	4	2.162162	0.044	<i>ITGA3, DKK1, CYR61, RNF165</i>
GOTERM_BP_FAT	GO:0048666~neuron development	16	8.648649	0.045	<i>BCL11A, ITGA3, PLK2, NTN4, NRXN3, NRG1, RORB, BMP7, ROBO1, LRFN5, VANGL2, FLRT3, ID2, CHN1, SLTRK6, RNF165</i>
GOTERM_BP_FAT	GO:1905207~regulation of cardiocyte differentiation	3	1.621622	0.047	<i>AKAP6, NRG1, DKK1</i>
GOTERM_BP_FAT	GO:0051410~detoxification of nitrogen compound	2	1.081081	0.047	<i>GSTM2, GSTM1</i>
GOTERM_BP_FAT	GO:0044272~sulfur compound biosynthetic process	6	3.243243	0.047	<i>GSTM2, GSTM1, VANGL2, ACOT11, ACSF2, GSTM5</i>
GOTERM_BP_FAT	GO:0030182~neuron differentiation	19	10.27027	0.047	<i>BCL11A, ITGA3, PLK2, NTN4, DIXDC1, NRXN3, NRG1, RORB, BMP7, DKK1, ROBO1, LRFN5, VANGL2, FLRT3, ID2, DUOX1, CHN1, SLTRK6, RNF165</i>
GOTERM_BP_FAT	GO:0007166~cell surface receptor signaling pathway	35	18.91892	0.048	<i>IRS1, HFE, ARHGEF28, DIXDC1, CYR61, EDA2R, ROBO1, DUOX1, FLRT3, CHN1, ATP6V0A4, ITGB6, GRIA3, TRPM3, ATP6V1C2, EVC2, KL, DST, CAV2, ITGA3, ITGA2, TNFRSF10C, NRG1, KIF7, BMP7, DKK1, PRLR, DNMT1, IL17RB, KRT19, VANGL2, COL4A5, SPRY1, ROR1, RNF165</i>
GOTERM_BP_FAT	GO:0048568~embryonic organ development	9	4.864865	0.048	<i>OSR2, KRT19, VANGL2, NRK, ID2, FOXI1, SLTRK6, BMP7, CYR61</i>
GOTERM_BP_FAT	GO:0045778~positive regulation of ossification	4	2.162162	0.049	<i>KL, OSR2, BMP7, CYR61</i>
GOTERM_BP_FAT	GO:0030501~positive regulation of bone mineralization	3	1.621622	0.049	<i>KL, OSR2, BMP7</i>
GOTERM_BP_FAT	GO:0055067~monovalent inorganic cation homeostasis	5	2.702703	0.049	<i>SCNN1G, RAB38, ATP6V0A4, CYP4F12, ATP6V1C2</i>
GOTERM_BP_FAT	GO:0048584~positive regulation of response to stimulus	28	15.13514	0.049	<i>IRS1, HFE, ARHGEF28, DIXDC1, CYR61, EDA2R, ROBO1, CHN1, S100A14, LMCD1, ATP6V1C2, KL, GSTM2, SCARA3, CAV2, LMO3, PLK2, ITGA2, NRG1, KIF7, BMP7, PRLR, IL17RB, VANGL2, POLR3G, AJUBA, RNF165</i>