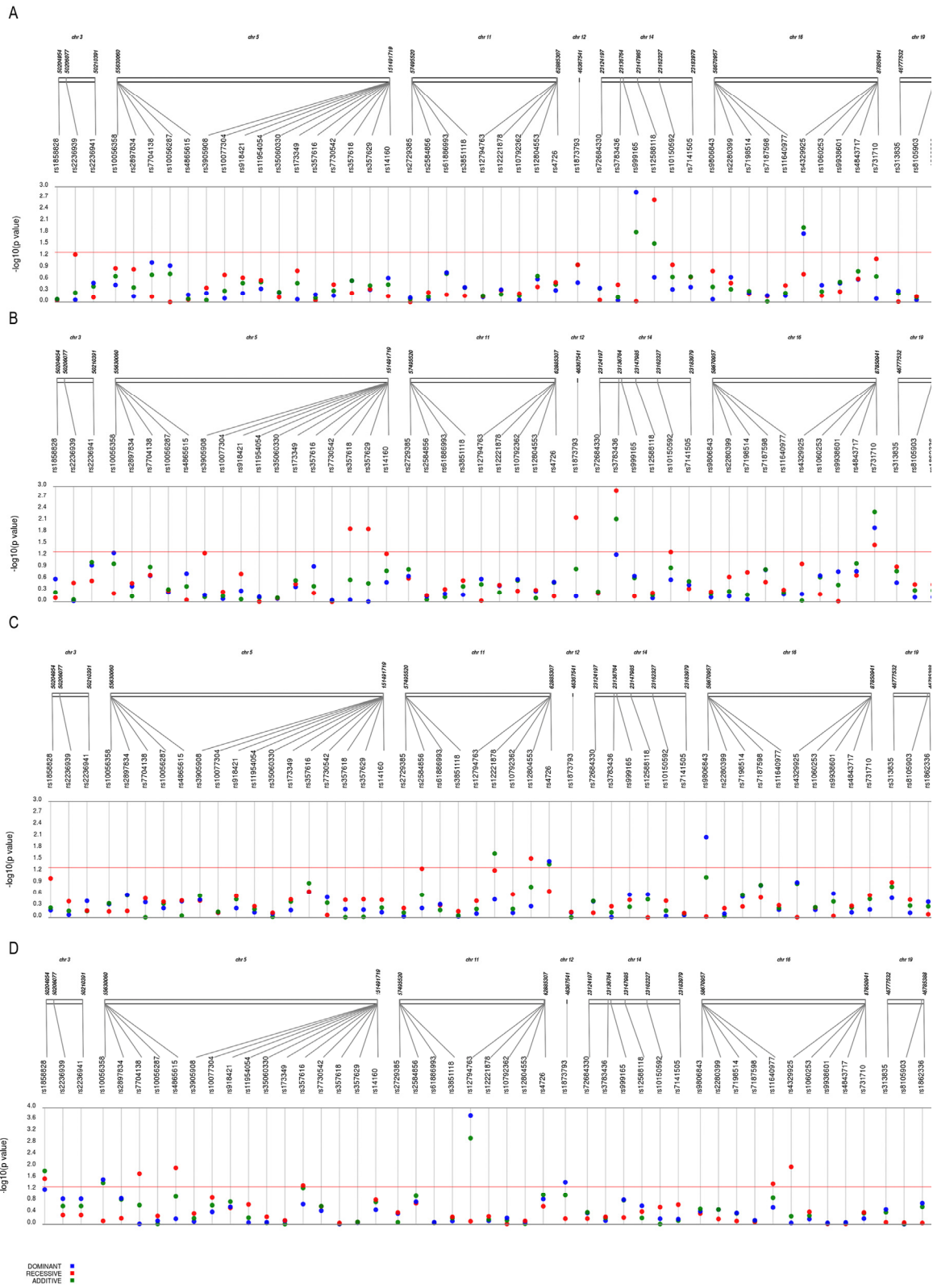


SUPPLEMENTARY MATERIAL



Supplementary Figure S1. Schematic representation of the association results with Hand Grip performance, Activity of Daily Living scores, Walking Time scores, and longevity (D).

Supplementary Table 1. Description of the 58 SNPs initially selected.

Gene/ Protein	Chr	SNP	Position (GRCh38.p7)	Mutation type (Ensembl)	Major/Minor allele (Ensembl)	MAF
<i>SLC38A3</i> / SNAT3	3	rs1858828	50204954	5'near gene variant	G/T	0.44
	3	rs2236939	50206077	Intron variant	G/T	0.11
	3	rs2236941	50210391	Intron variant	C/T	0.32
<i>SLC36A1</i> / PAT1	5	rs3905908	151445768	5'near gene variant	T/C	0.22
	5	rs10077304	151452320	Intron variant	T/C	0.16
	5	rs918421	151452428	5'UTR variant	A/G	0.46
	5	rs11954054	151456584	Intron variant	A/G	0.23
	5	rs35060330	151458524	Intron variant	C/T	0.33
	5	rs17112008	151458557	Intron variant	C/A	0.33
	5	rs173349	151462020	Intron variant	T/C	0.47
	5	rs357616	151465808	Intron variant	T/C	0.40
	5	rs7730542	151466243	Intron variant	A/G	0.12
	5	rs357618	151467051	Intron variant	A/G	0.39
	5	rs357629	151471385	Intron variant	A/G	0.39
	5	rs14160	151491719	3'UTR variant	T/C	0.28
	5	rs1175	151492234	3'UTR variant	C/T	0.20
	<i>SLC38A9</i> / SNAT9	5	rs10056358	55630060	Intron variant	A/T
5		rs2897834	55642751	Intron variant	C/A	0.13
5		rs7704138	55648434	Intron variant	C/T	0.34
5		rs10056287	55658650	Intron variant	T/C	0.19
5		rs4865615	55664845	Missense variant	C/G	0.34
5		rs7735053	55667626	Intron variant	T/C	0.34
5		rs11749532	55706437	Intron variant	G/A	0.30
5		rs7736177	55708116	Intron variant	G/A	0.31
<i>SLC3A2</i> / CD98	11	rs12794763	62858040	Intron variant	T/G	0.15
	11	rs12221878	62858559	Intron variant	C/G	0.04
	11	rs10792362	62873879	Intron variant	T/C	0.43
	11	rs12804553	62876155	Intron variant	G/T	0.28
	11	rs4726	62885307	Exon Synonymous variant	C/T	0.25
	11	rs2282477	62889032	3'near gene variant	T/C	0.23
<i>SLC43A1</i> / LAT3	11	rs2729385	57495520	Intron variant	G/A	0.33
	11	rs2584856	57499839	Intron variant	C/A	0.23
	11	rs61886993	57511522	Intron variant	C/G	0.08
	11	rs3851118	57513478	Intron variant	T/C	0.35

<i>SLC7A8/</i> <i>LAT2</i>	14	rs72684330	23124197	3'near gene variant	T/A	0.12
	14	rs17794251	23124233	3'near gene variant	C/T	0.27
	14	rs3783436	23136764	Intron variant	T/C	0.34
	14	rs999165	23138657	Intron variant	T/A	0.27
	14	rs12588118	23147985	Intron variant	C/G	0.23
	14	rs10150592	23162327	Intron variant	C/A	0.17
	14	rs7141505	23183979	5'near gene variant	C/A	0.29
<i>SLC7A5/</i> <i>LAT1</i>	16	rs4329925	87828401	3'near gene variant	T/C	0.14
	16	rs1060253	87832532	3'near gene variant	G/C	0.26
	16	rs9938601	87837658	Intron variant	A/G	0.42
	16	rs4843717	87844559	Intron variant	T/C	0.27
	16	rs731710	87850941	Intron variant	A/G	0.41
	16	rs7193392	87854200	Intron variant	G/A	0.39
<i>SLC38A7/</i> <i>SNAT7</i>	16	rs9806843	58670957	Intron variant	A/G	0.42
	16	rs2280399	58677628	Intron variant	G/T	0.11
	16	rs7198514	58680599	Intron variant	C/T	0.20
	16	rs7187598	58681490	Non coding exon variant	T/C	0.41
	16	rs11640977	58681712	Splice region variant	T/C	0.14
	16	rs8058969	58685026	5'UTR variant	G/A	0.49
<i>SLC1A5/</i> <i>ASCT2</i>	19	rs313835	46777532	Intron variant	C/T	0.28
	19	rs8105903	46784893	Intron variant	A/C	0.47
	19	rs1862336	46785388	5'UTR variant	T/C	0.24

Abbreviations: MAF (Minor Allele Frequency).

MAF refers to the European population as retrieved by Ensembl (www.ensembl.org)

Table S2. Summary of functional annotation of the phenotype-associated SNPs.

GENE	SNP	SNPs in LD ($r^2 \geq 0.8$)	HaploReg v4.1							dbSNP func annot	RegulomeDB Score
			Promoter histone marks	Enhancer histone marks	DNase	Proteins bound	Motifs changed	GRASP QTL hits	Selecte d eQTL hits		
<i>SLC3A2</i> CD98	rs12804553	4	SKIN	6 tissues	SKIN, PLCNT		4 altered motifs		8 hits	intronic	3a less likely to affect binding
	rs4726	8		15 tissues	7 tissues	POL2, POL24H8	Maf		7 hits	synonymous	4 Minimal binding evidence
	rs12794763	none	8 tissues	12 tissues			Foxl1, Pou1f1	1 hits	4hits	intronic	3a less likely to affect binding
<i>SLC7A5</i> LAT1	rs4329925	31		12 tissues	13 tissues	POL24H8	Ets,RBP-Jkappa		1 hits	1.6kb 3' of SLC7A5	2b likely to affect binding
	rs731710	4		19 tissues	7 tissues		ATF3,ATF6,RFX5		3 hits	intronic	2b likely to affect binding
<i>SLC7A8</i> LAT2	rs999165	none		8 tissues	GI,BLD		HDAC2,PRDM1,TATA			intronic	No Data
	rs12588118	8		13 tissues	4 tissues		4 altered motifs			intronic	4 Minimal binding evidence
	rs3783436	5		6 tissues	HRT,LIV		NF-Y	5 hits	5 hits	intronic	5 minimal binding evidence
<i>SLC36A1</i> PAT1	rs357618	22					Egr-1		46 hits	intronic	6
	rs357629	22		FAT, SKIN	SKIN,SKN		Nkx3,Pou2f2	1 hits	46 hits	intronic	1f Likely to affect binding and linked to expression of a gene target
<i>SLC38A2</i> SNAT2	rs1873793	9	HRT	17 tissues	10 tissues	4 bound proteins	6 altered motifs	1 hits	7 hits	intronic	2b likely to affect binding
	rs1858828	14	10 tissues	17 tissues	5 tissues		BDP1,Ets, RXRA		41 hits	291bp 5' of SLC38A3	4 Minimal binding evidence
<i>SLC38A7</i> SNAT7	rs9806843	2					FAC1,NF-I	1 hits	3 hits	intronic	4 Minimal binding evidence

SLC38A 9 SNAT9	rs4865615	324		FAT, MUS			4 altered motifs		3 hits	missense	6 Minimal binding evidence
	rs7704138	318						1 hits	4 hits	intronic	5 minimal binding evidence
	rs10056358	11			4 tissues		Egr1, SETDB1	2 hits		intronic	4 Minimal binding evidence

Abbreviations: Promoter/Enhancer histone marks, regulatory chromatin states based on ENCODE and Epigenomics Roadmap data; DNase, DNase hypersensitivity based on Epigenomics Roadmap data; Proteins bound, proteins bound by chromatin immunoprecipitation based on Epigenomics Roadmap data; Motifs changed, altered regulatory motifs; GRASP QTL hits, quantitative trait loci based on GRASP (Genome-Wide Repository of Associations Between SNPs and Phenotypes); selected eQTL hits, expression quantitative trait loci based on the Genotype-Tissue Expression (GTEx) analysis.

Scores indicate the following degrees of evidence: Score 1a, eQTL + TF binding + matched TF motif + matched DNase Footprint + DNase peak; Score 2b, TF binding + any motif + DNase Footprint + DNase peak; Score 3a, TF binding + any motif + DNase peak; Score 4, TF binding + DNase peak; Score 5, TF binding or DNase peak; Score 6, other; "No data" indicates that RegulomeDB holds no information about the given SNP, meaning there currently exists no evidence to suggest that the SNP has a regulatory function.