

Detection and replication of epistasis influencing transcription in humans

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Abstract

Epistasis is the phenomenon whereby one polymorphism's effect on a trait depends on other polymorphisms present in the genome. The extent to which epistasis influences complex traits¹ and contributes to their variation^{2,3} is a fundamental question in evolution and human genetics. Though epistasis has been demonstrated in artificial gene manipulation studies in model organisms,^{4,5} and some examples have been reported in other species,⁶ few convincing examples exist for epistasis amongst natural polymorphisms in human traits.^{7,8} Its absence from empirical findings may simply be due to its low incidence in the genetic control of complex traits,^{2,3} but an alternative view is that it has previously been too technically challenging to detect due to statistical power and computational issues.⁹ Here we show that, using advanced computation techniques¹⁰ and a gene expression study design, many instances of epistasis are found between common single nucleotide polymorphisms (SNPs). In a cohort of 846 individuals with data on 7339 gene expression levels in whole blood, we found 501 significant pairwise epistatic interactions between common SNPs acting on the expression levels of 238 genes ($p < 2.91 \times 10^{-16}$). We tested the discovery interactions for replication in two independent data sets.^{11,12} Three hundred and forty-five interactions had replication interaction p -values that were more extreme than the 2.5% confidence interval of the distribution under the null hypothesis of no epistasis, with 30 significant at a conservative $p < 0.05$ Bonferroni level. There was evidence of functional enrichment for the interacting SNPs, for instance 44 of the genetic interactions are located within 2Mb of regions of known intra-cellular chromosome interactions¹³ ($p = 1.8 \times 10^{-10}$). Epistatic networks of three SNPs or more influence the expression levels of 129 genes, whereby one *cis*-acting SNP is modulated by several *trans*-acting SNPs. For example MBNL1 is influenced by an additive effect at rs13069559 which itself is masked by *trans*-SNPs on 14 different chromosomes, with nearly identical genotype-phenotype (GP) maps for each *cis-trans* interaction. This study presents the first evidence for multiple instances of epistatic genetic effects emerging from natural genetic variation in humans.

1 Main text

In the genetic analysis of complex traits it is usual for SNP effects to be estimated using an additive model where they are assumed to contribute independently and cumulatively to the mean of a trait. This framework has been successful in identifying thousands of associations,¹⁴ but to date there is little empirical exploration of the role that epistasis plays in the architecture of complex traits in humans,^{7,8} though its contribution to phenotypic variance is frequently the subject of debate.¹⁻³ Outside the prism of human association studies there is evidence for epistasis, not only at the molecular scale from artificially induced mutations⁴ but also at the evolutionary scale in fitness adaptation¹⁵ and speciation.¹⁶

Methods are now available to overcome the computational problems involved in searching for epistasis, but its detection still remains problematic due to re-

duced statistical power. For example increased dependence on linkage disequilibrium (LD) between causal SNPs and observed SNPs,^{17,18} increased model complexity in fitting interaction terms,¹⁹ and more extreme significance thresholds to account for increased multiple testing⁹ all make it more difficult to detect epistasis in comparison to additive effects. When genetic effect sizes are small, as is expected in most complex traits of interest,¹⁴ the power to detect epistasis diminishes rapidly. There are two simple ways to overcome this problem. One is by using extremely large sample sizes;²⁰ another is by analysing traits that are likely to have large effect sizes. Because our focus was to ascertain the extent to which instances of epistasis occur amongst natural genetic variation we designed a study around the latter approach and searched for epistatic genetic effects that influence gene expression levels. Transcription levels can be measured for thousands of genes. These traits are largely heritable but on average less polygenic than high level phenotypes,²¹ thus it is expected that many genetic effects will be relatively large, maximising the chance at detecting epistasis, should it exist.

In our discovery dataset (Brisbane Systems Genetics Study, BSGS²²) of 846 individuals genotyped at 528,509 SNPs, we exhaustively tested every pair of SNPs for genetic interactions against each of 7339 expression traits in whole blood. After stringent filtering and multiple testing correction (Methods) we identified 501 putative genetic interactions influencing the expression levels of 238 genes. Of the 501 discovery interactions, 434 had available data and passed filtering (Methods) in two independent replication datasets, Fehrman¹² and the Estonian Genomics Centre University of Tartu (EGCUT),¹¹ in which we saw convincing evidence for replication. We used the summary statistics from the replication datasets to perform a meta analysis to obtain an independent p -value for the putative interactions, and 30 were significant after applying a Bonferroni correction for multiple testing (Table 1). These significant interactions exhibited remarkable similarity in GP maps between all three datasets (Figure 2).

In addition, we observed that 316 of the remaining 404 discovery SNPs had replication interaction p -values exceeding the one-tailed 2.5% confidence interval under the null distribution of no effects ($p \ll 1.0 \times 10^{-16}$, Figure 3 and Supplementary Figure S1). The congruence of the epistatic networks in discovery and replication datasets is shown in Figure 1, demonstrating that these complex genetic patterns are common even across independent datasets. A further replication was attempted using the Centre for Health Discovery and Wellbeing (CHDWB) dataset,²³ but only 185 of the SNP pairs passed filtering because the sample size was small ($n = 139$), and likely due to insufficient power we found no evidence for replication. It should be noted that although it is a necessary step to establish the veracity of the signals from the discovery set, replication of epistasis is theoretically difficult because the dependence on LD between observed SNPs and causal variants is up to four orders of magnitude higher than it is for independent additive effects.^{17,18} Therefore these results are encouraging with regards to the detection and replication of epistasis.

Though seldom the focus of association studies, SNPs with known main effects are often tested for additive \times additive genetic interactions,⁹ but our analysis shows that this is unlikely to be the most effective strategy for its detec-

tion. The majority of our discovery interactions comprised of one SNP that was significantly associated with the gene expression level in the discovery dataset, and one SNP that had no previous association²¹ (439 out of 501, Methods). Only nine interactions were between SNPs that both had known main effects while 64 were between SNPs that had no known main effects. Additionally, we observed that the largest epistatic variance component for the 501 interactions was equally divided amongst additive \times additive, additive \times dominance, dominance \times additive and dominance \times dominance ($p = 0.22$ for departure from expectation). This is not surprising because the patterns of epistasis used for statistical decomposition are not designed to resemble biological function.²⁴

Of the discovery interactions, 47 were *cis-cis* acting (both SNPs were on the same chromosome as the expression gene), 441 were *cis-trans*-acting, and 13 were *trans-trans*-acting. We observed a wide range of significant GP maps (Figure 2) but the most common pattern of epistasis that we detected involved a *trans*-SNP masking the effect of an additive *cis*-SNP. For example, MBNL1 (involved in RNA modification and regulation of splicing²⁵) has a *cis* effect at rs13069559 which in turn is controlled by 13 *trans*-SNPs and one *cis*-SNP that each exhibit a masking pattern, such that when the *trans*-SNP is homozygous for the masking allele the decreasing allele of the *cis*-SNP no longer has an effect (Supplementary Figure S5). Each of these interactions have evidence for replication in at least one dataset and six are significant at the Bonferroni level (Supplementary Figure S2). We see similar epistatic networks involving multiple *trans*-acting SNPs for other gene expression levels too, for example TMEM149 (Supplementary Figure S6), NAPRT1 (Supplementary Figure S7), TRAPPC5 (Supplementary Figure S8), and CAST (Supplementary Figure S9).

In total the 501 interactions comprised 781 unique SNPs, which we analysed for functional enrichment (Methods). We tested the SNPs for cell-type specific overlap with transcriptionally active chromatin regions, tagged by histone-3-lysine-4,3-methylation (H3K4me3) chromatin marks, in 34 cell types²⁶ (Supplementary Figure S4). There was significant enrichment for *cis*-acting SNPs in haematopoietic cell types only ($p < 1 \times 10^{-4}$ for the three tissues with the strongest enrichment after adjusting for multiple testing). However *trans*-acting SNPs did not show any tissue specific enrichment ($p > 0.1$ for all tissues). This difference between *cis* and *trans* SNPs suggests different roles in which epistasis might arise where the *cis*-SNPs provide tissue specificity in these interactions. There is also strong enrichment for SNPs to be localised in enhancer regions,²⁷ consistent for both *cis* and *trans* SNPs ($p < 1 \times 10^{-6}$).

We also demonstrate spacial organisation of interacting loci suggesting a mechanism by which biological function can lead to epistatic genetic variance. It has been shown that different chromosomal regions spatially colocalise in the cell through chromatin interactions.¹³ We cross-referenced our epistatic SNPs with a map of chromosome interacting regions ($n = 96, 139$) in K562 blood cell lines²⁸ (Methods) and found that 44 epistatic interactions mapped to within 2Mb ($p < 1.8 \times 10^{-10}$), (Supplementary Figure S10). Interaction of distant loci may occur through physical proximity in transcriptional factories that organise across different chromosome regions and can regulate transcription of related

genes.^{29,30}

Though we present many instances of epistasis, quantifying its relative importance to complex traits in humans remains an open question. In this study we are able to identify 238 gene expression traits with at least one significant interaction given our experiment-wide threshold. How does this compare to the number of traits influenced by additive effects? The BSGS dataset has been previously analysed for additive effects at all expression traits,²² and if we take all the additive eQTLs that were significant at the epistatic threshold of $p < 2.91 \times 10^{-16}$ we find that 453 gene expression levels out of the 7339 analysed had at least one significant expression quantitative trait locus (eQTL). Therefore it can be argued that the number of instances of detectable epistasis are substantial.

However in terms of their contribution to complex traits a more important metric might be the proportion of the variance that the epistatic loci explain.² Ideally one would approach this question from a whole genome perspective³¹ but this is intractable for non-additive variance components. Nevertheless, some inference can be made from the ascertained effects in these analyses and it is evident that additive variance is overall a larger component than epistatic variance, as has been argued previously.^{2,3} Taking the additive effects detected in Powell *et al* (2012) at the $p < 2.91 \times 10^{-16}$ threshold, we calculate that on average they explain 1.73% of the phenotypic variance of each of the 7339 probes. By contrast, the epistatic variance from the interacting SNPs detected in this study on average explain 0.25% of phenotypic variance, approximately seven times lower than the additive variance (Methods). There are at least three caveats to this comparison. Firstly, the ratio of additive to epistatic variance may differ at different effect sizes. Secondly, the power of a 1 *d.f.* test exceeds that of an 8 *d.f.* test. And thirdly, the non-additive variance at causal variants is expected to be underestimated by observed SNPs in comparison to estimates for additive variance, due to differences in the rate of decay of the estimate of the genetic variance of the causal SNPs as LD decreases with the observed SNPs.

Overall, we have demonstrated that it is possible to identify and replicate epistasis in complex traits amongst common human variants. The functional analysis of the significant epistatic loci suggests that there are a large number of possible mechanisms that can lead to non-additive genetic variation. Further research into such epistatic effects may provide a useful portal to understanding molecular mechanisms and complex trait variation with greater clarity. With computational techniques and data now widely available the search for epistasis in larger datasets for traits of broader interest is warranted.

1.1 Methods Summary

We searched for pairwise epistasis exhaustively in the BSGS discovery dataset,²² which comprises 846 individuals who are genotyped at 528,509 autosomal SNPs and who have gene expression levels measured in whole blood samples for 7,339 probes representing 6,158 RefSeq genes. Recent hardware and software¹⁰ ad-

vances made it possible to perform the 1.03×10^{15} statistical tests to complete this analysis. We used permutation analysis³² to calculate an experiment-wide significance threshold of $T_e = 2.91 \times 10^{-16}$ at the 5% family-wise error rate (FWER). SNP pairs were modelled for full genetic effects, including marginal additive and dominance at both SNPs plus four interaction terms. Though we could have used a less complex model to improve statistical efficiency, we deemed it important to be agnostic about the type of epistasis that might exist, and therefore chose not to over-parameterise the test.^{18,19} Because there are many large marginal effects present in these data it was necessary to perform several filtering steps to exclude SNP pairs that were significant due to marginal effects alone. All SNP pairs with LD $r^2 > 0.1$ and $D'^2 > 0.1$ were removed to minimise the possibility of haplotype effects. All SNP pairs were required to have at least five data points in all nine genotype classes. If multiple SNP pairs were present on the same chromosomes for a particular expression trait then only the sentinel SNP pair was retained. Finally, a nested test contrasting the full genetic model against the marginal additive and dominance model was performed for each remaining SNP pair (Methods), resulting in 501 significant interactions after Bonferroni correction for multiple testing of the filtered SNPs. The significant SNP pairs were carried forward for replication in two independent datasets that used the same expression assays for analysing transcription in whole blood, the Fehrman dataset¹² ($n = 1240$) and the Estonian Genome Centre University of the University of Tartu (EGCUT) dataset¹¹ ($n = 891$). Of these, 434 passed filtering in both replication datasets. A meta analysis on the interaction p -values from each replication dataset was performed to provide an overall replication statistic for each putative interaction.

1.2 Acknowledgements

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2 Tables

Table 1: Epistatic interactions significant at the Bonferroni level in two replication sets

	Gene (chr.)	SNP 1 (chr.)	SNP 2 (chr.)	BSGS ²	Fehrmann ³	EGCUT ³	Meta ⁴
1	ADK (10)	rs2395095 (10)	rs10824092 (10)	6.69 ¹	18.33 ¹	21.21 ¹	39.82 ¹
2	ATP13A1 (19)	rs4284750 (19)	rs873870 (19)	5.30	12.18	3.25	14.23
3	C21ORF57 (21)	rs9978658 (21)	rs11701361 (21)	9.42	6.08	16.36	21.67
4	CSTB (21)	rs9979356 (21)	rs3761385 (21)	11.99	25.20	16.72	42.27
5	CTSC (11)	rs7930237 (11)	rs556895 (11)	7.16	18.76	15.06	33.53
6	FN3KRP (17)	rs898095 (17)	rs9892064 (17)	16.16	28.24	29.39	59.95
7	GAA (17)	rs11150847 (17)	rs12602462 (17)	13.91	19.98	12.99	32.60
8	HNRPH1 (5)	rs6894268 (5)	rs4700810 (5)	15.38	8.55	3.01	10.37
9	LAX1 (1)	rs1891432 (1)	rs10900520 (1)	19.16	18.60	11.22	29.24
10	MBNL1 (3)	rs16864367 (3)	rs13079208 (3)	13.49	16.25	24.74	41.56
11	MBNL1 (3)	rs7710738 (5)	rs13069559 (3)	7.92	2.55	7.89	9.28
12	MBNL1 (3)	rs2030926 (6)	rs13069559 (3)	7.10	0.91	5.80	5.53
13	MBNL1 (3)	rs2614467 (14)	rs13069559 (3)	5.74	4.13	2.22	5.30
14	MBNL1 (3)	rs218671 (17)	rs13069559 (3)	7.63	0.62	5.82	5.23
15	MBNL1 (3)	rs11981513 (7)	rs13069559 (3)	7.71	0.43	5.36	4.58
16	MBP (18)	rs8092433 (18)	rs4890876 (18)	5.40	7.06	21.91	28.73
17	NAPRT1 (8)	rs2123758 (8)	rs3889129 (8)	8.45	15.12	16.08	30.77
18	NCL (2)	rs7563453 (2)	rs4973397 (2)	7.31	7.51	6.33	12.70
19	PRMT2 (21)	rs2839372 (21)	rs11701058 (21)	4.81	0.69	4.47	4.06
20	RPL13 (16)	rs352935 (16)	rs2965817 (16)	4.98	3.79	14.41	17.24
21	SNORD14A (11)	rs2634462 (11)	rs6486334 (11)	7.31	13.11	10.96	23.22
22	TMEM149 (19)	rs807491 (19)	rs7254601 (19)	12.16	81.55	45.78	145.78
23	TMEM149 (19)	rs8106959 (19)	rs6926382 (6)	5.80	3.06	8.80	10.72
24	TMEM149 (19)	rs8106959 (19)	rs914940 (1)	6.22	3.36	6.96	9.20
25	TMEM149 (19)	rs8106959 (19)	rs2351458 (4)	7.30	0.04	9.61	8.00
26	TMEM149 (19)	rs8106959 (19)	rs6718480 (2)	8.55	3.31	5.15	7.36
27	TMEM149 (19)	rs8106959 (19)	rs1843357 (8)	6.21	3.72	3.33	6.00
28	TMEM149 (19)	rs8106959 (19)	rs9509428 (13)	9.44	0.10	5.75	4.47
29	TRA2A (7)	rs7776572 (7)	rs11770192 (7)	8.23	3.19	1.89	4.09
30	VASP (19)	rs1264226 (19)	rs2276470 (19)	5.09	0.94	5.14	4.95

¹ $-\log_{10} p$ -values for 4 *d.f.* interaction tests

² Discovery dataset

³ Independent replication dataset

⁴ Meta analysis of interaction terms between replication datasets only

3 Figures

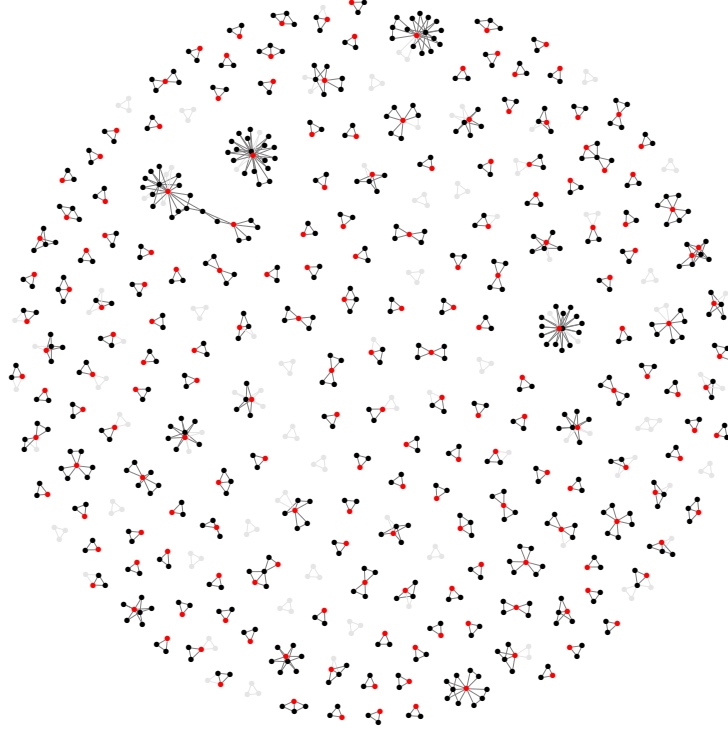


Figure 1: **Discovery and replication of epistatic networks** All 434 putative genetic interactions (edges) with data common to discovery and replication sets is shown, where black nodes represent SNPs and red nodes represent traits (gene expression probes). Three hundred and forty-five interactions had p -values exceeding the 2.5% confidence interval following meta analysis of the replication data, but the remaining 89 interactions that did not replicate are depicted in grey. It is evident that a large proportion of the complex networks identified in the discovery set also exist in independent populations.

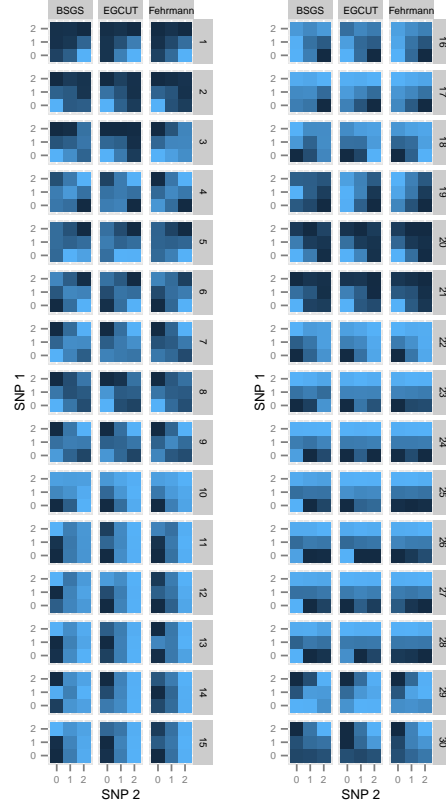


Figure 2: Replication of GP maps in two independent populations The GP maps for each epistatic interaction that is significant at the Bonferroni level in both replication datasets are shown. Each GP map consists of nine tiles where each tile represents the expression level for that two-locus genotype class. Phenotypes are for gene transcript levels (dark coloured tiles = low expression, light coloured tiles = high expression). Columns of GP maps are for each independent population. Rows of GP maps are for each of 30 significantly replicated interactions at the Bonferroni level, corresponding to the rows in Table 1. There is a clear trend of the GP maps replicating across all three datasets.

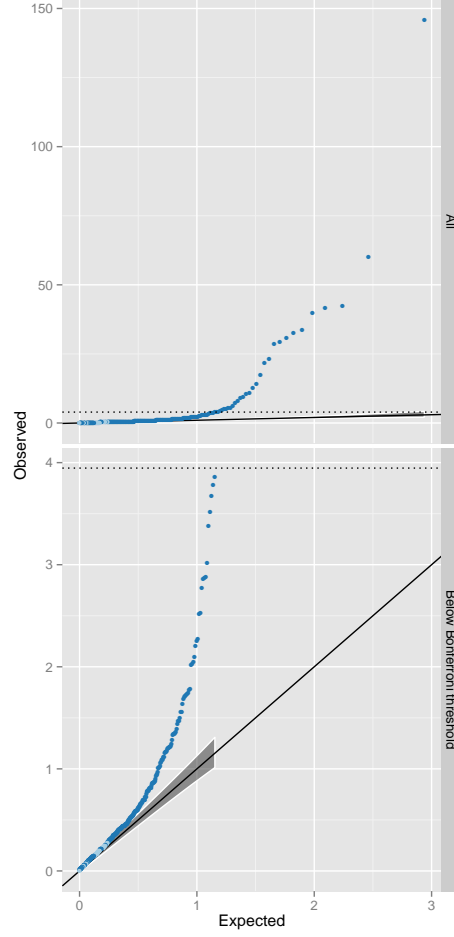


Figure 3: **Q-Q plots of interaction p -values from replication datasets** The top panel shows all 434 discovery SNPs that were tested for interactions. Observed p -values (y -axis, $-\log_{10}$ scale) are plotted against the expected p -values (x -axis, $-\log_{10}$ scale). The multiple testing correction threshold for significance following Bonferroni correction is denoted by a dotted line. The bottom panel shows the same data as the top panel but excluding the 30 interactions that were significant at the Bonferroni level in the replication datasets. The shaded grey area represents the 5% confidence interval for the expected distribution of p -values. Dark blue points represent p -values that exceed the confidence interval, light blue are within the confidence interval.

4 Supplementary Figures

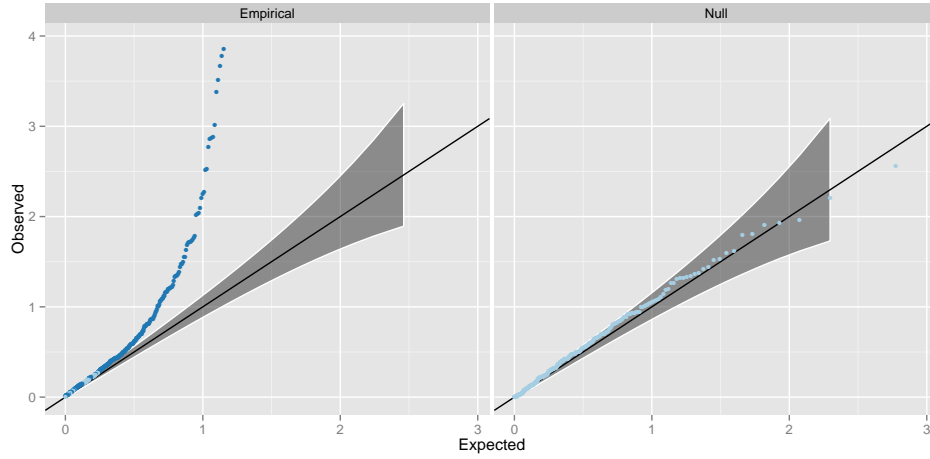


Figure S1: **Q-Q plots of interaction p -values from replication datasets, excluding the 30 points significant at the Bonferroni level** The right panel (Null) shows the interaction p -values from a meta analysis across two independent datasets on 434 randomly drawn SNP pairs. The left panel (Empirical) shows the interaction p -values from the 404 putative interactions that were not significant at the Bonferroni correction threshold. Dark blue points represent p -values that surpass the 2.5% FDR level, as in Figure 3.

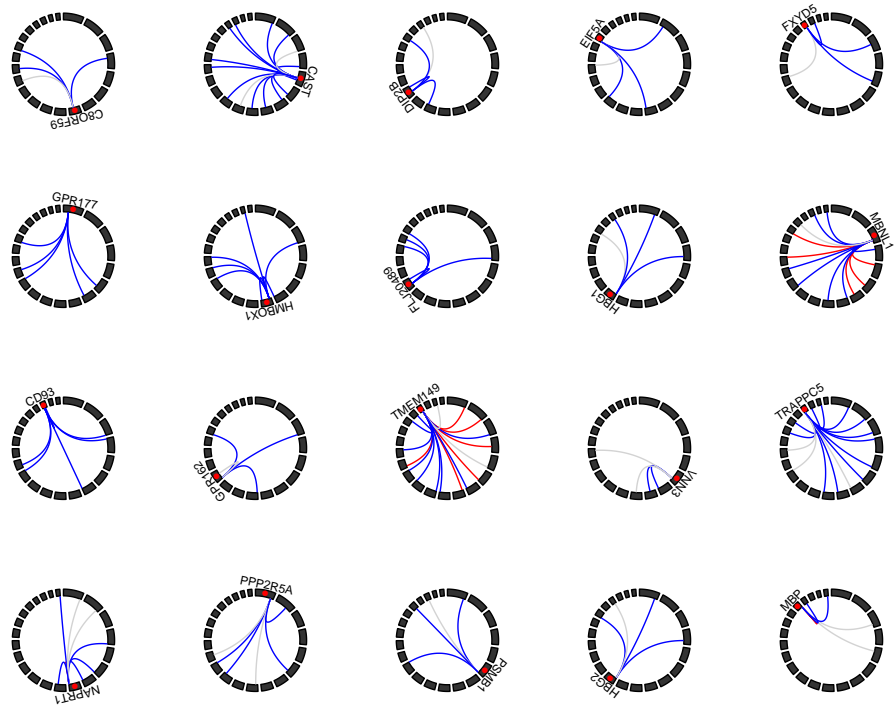


Figure S2: **Gene expression traits with four or more genetic interactions** Circle plots represent the genomic positions for SNPs (linking lines) and expression probes (red points). Chromosomes are represented by black blocks and ordered from 1 to 22 clockwise, starting from the top. Grey lines represent no evidence for replication, blue lines denote interactions that are outside the 97.5% confidence interval or the Q-Q plot (Figure 3), and red lines denote replication at the Bonferroni correction level. Most interactions are characterised as being *cis-trans* to the expression probe.

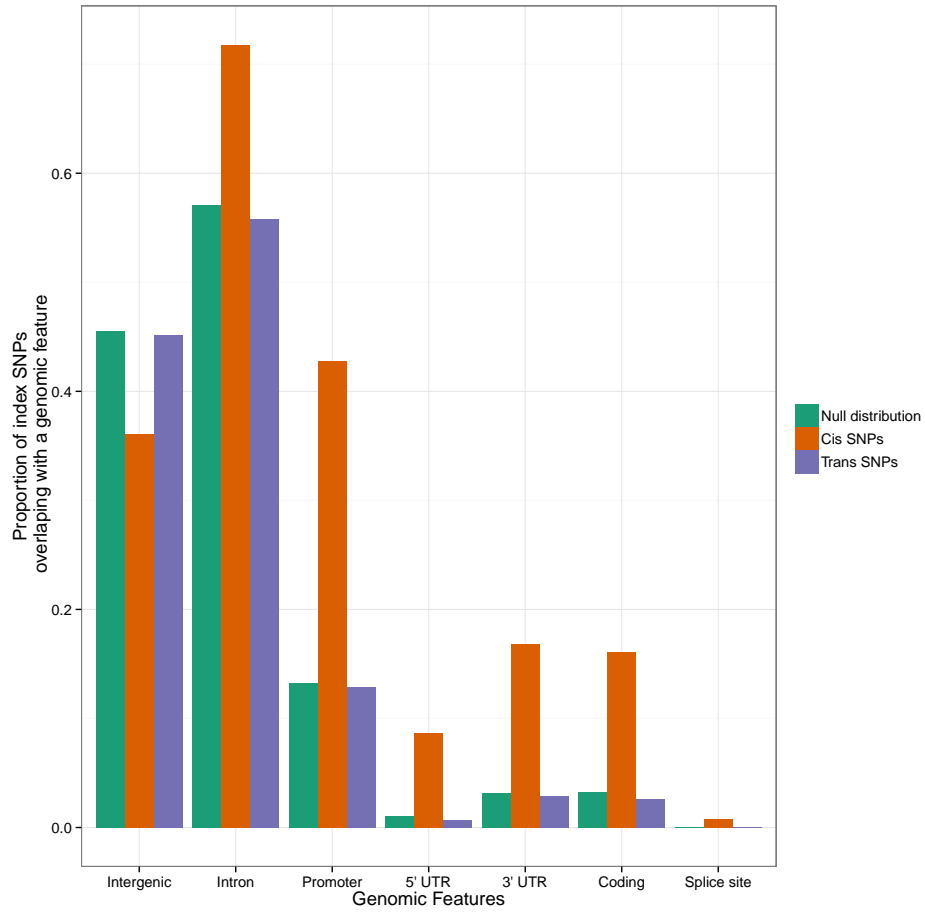


Figure S3: Location of SNPs relative to genomic features All SNPs within 1kb and $r^2 > 0.8$ of each *cis*- and *trans*-SNP were taken to find which genomic features (*x*-axis) were covered by the SNPs that compose the 501 significant interactions. Green bars represent the proportion (*y*-axis) of the 528,509 SNPs used in the analysis that fall within the range of the different genomic features. There is enrichment for *cis*-acting SNPs (red bars) in promotor regions, but *trans*-acting SNPs (blue bars) are not enriched for genomic features.

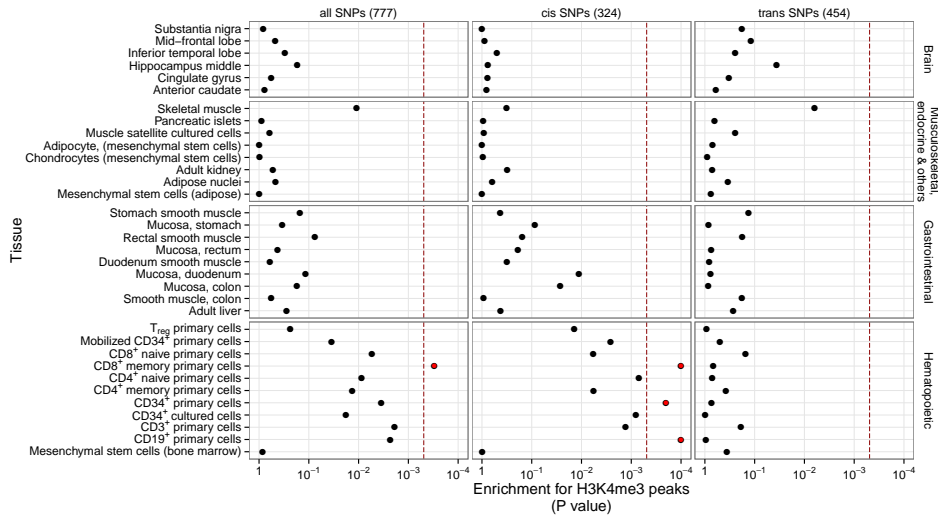


Figure S4: Tissue specific enrichment of SNPs in transcriptionally active regions The locations of transcriptional activity can be predicted by chromatin marks, assayed by H3K4me3.²⁶ Enrichment p -values are calculated using permutation analysis for 34 different cell types (y -axis) in four tissue types (Rows of boxes). There is enrichment for *cis*-acting SNPs in Haematopoietic tissue types only. *Trans*-acting SNPs have no tissue specificity.

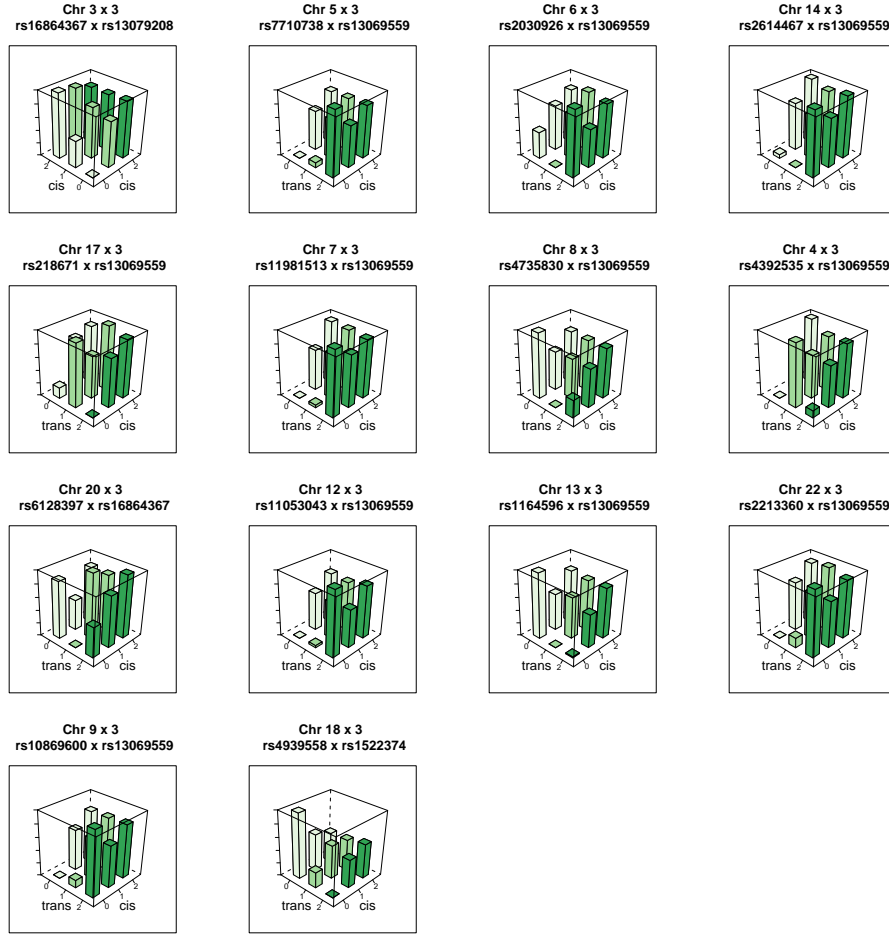


Figure S5: **Genotype-phenotype maps for 14 interactions influencing the expression of MBNL1** Each bar represents the mean phenotypic value for individuals in that genotype class. The rs13069559 SNP typically has a *cis*-additive decreasing effect on the expression of MBNL1, but in many of these interactions the *cis* effect is masked when the *trans* SNP is homozygous.

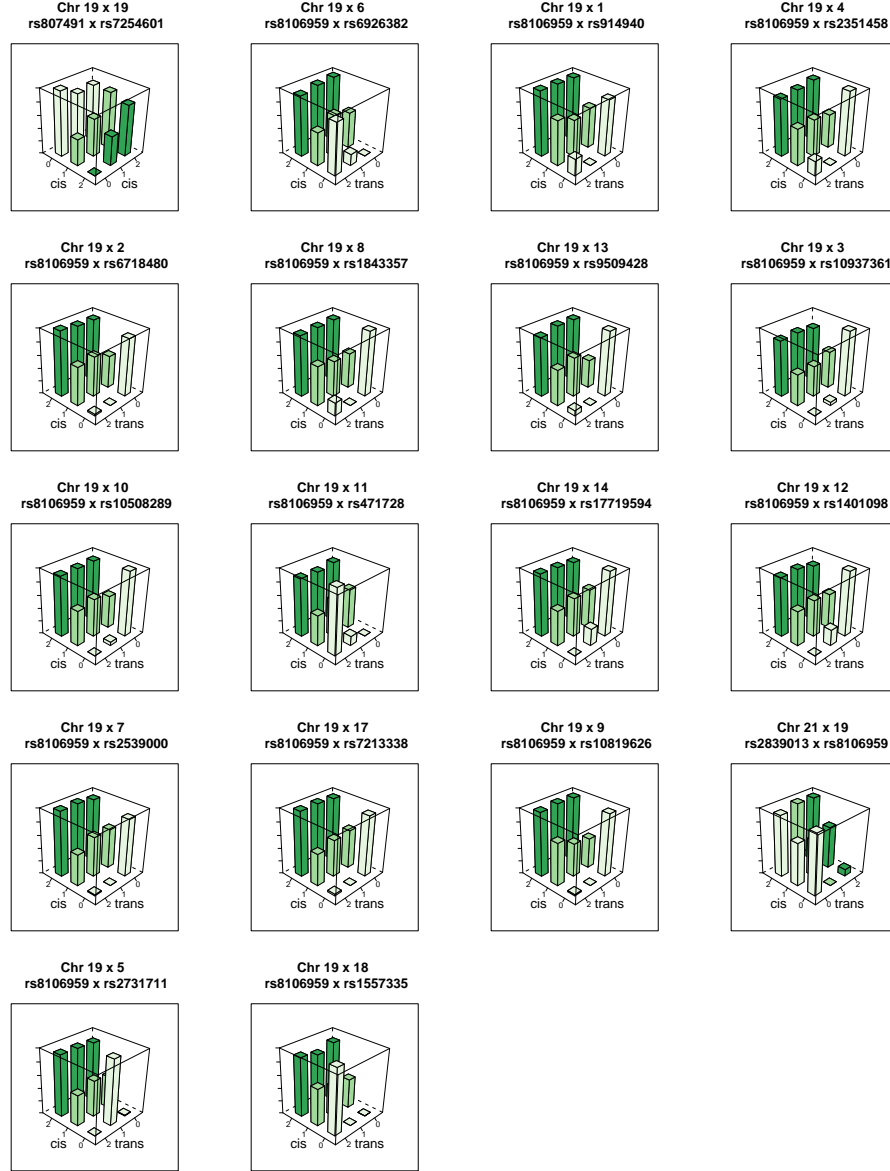


Figure S6: **Genotype-phenotype maps for 19 interactions influencing the expression of TMEM149** Each bar represents the mean phenotypic value for individuals in that genotype class. The rs13069559 SNP typically has a *cis*-additive decreasing effect on the expression of TMEM149, but in many of these interactions the *cis* effect is masked when the *trans* SNP is homozygous.

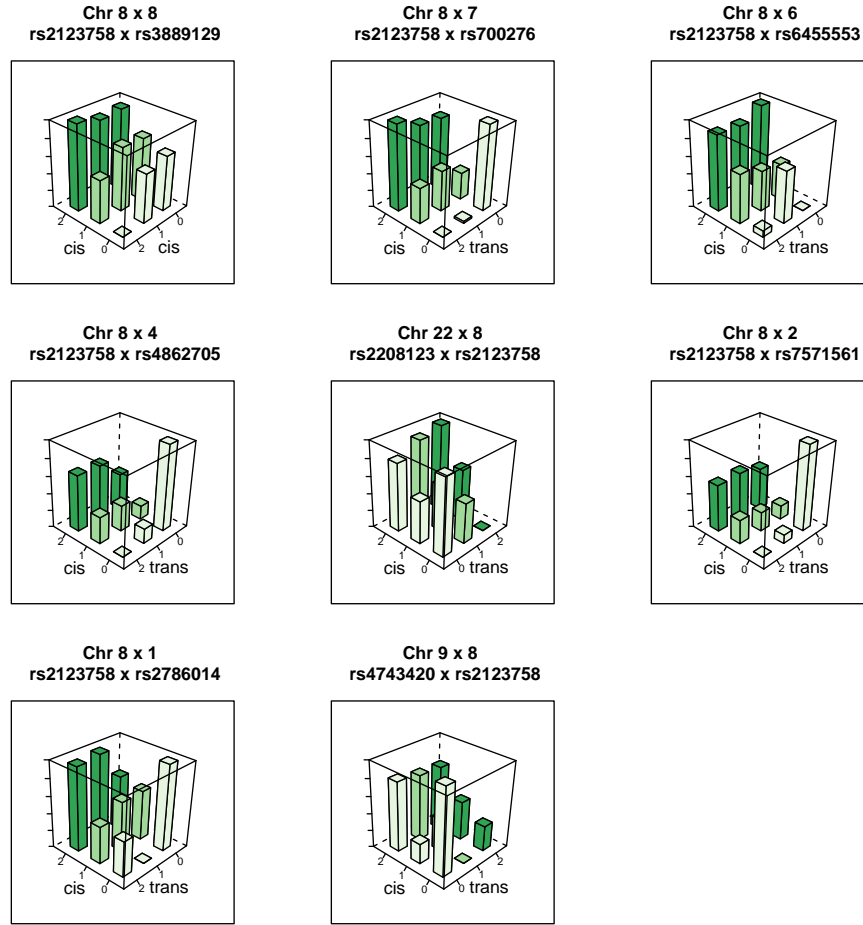


Figure S7: **Genotype-phenotype maps for 8 interactions influencing the expression of NAPRT1** Each bar represents the mean phenotypic value for individuals in that genotype class.

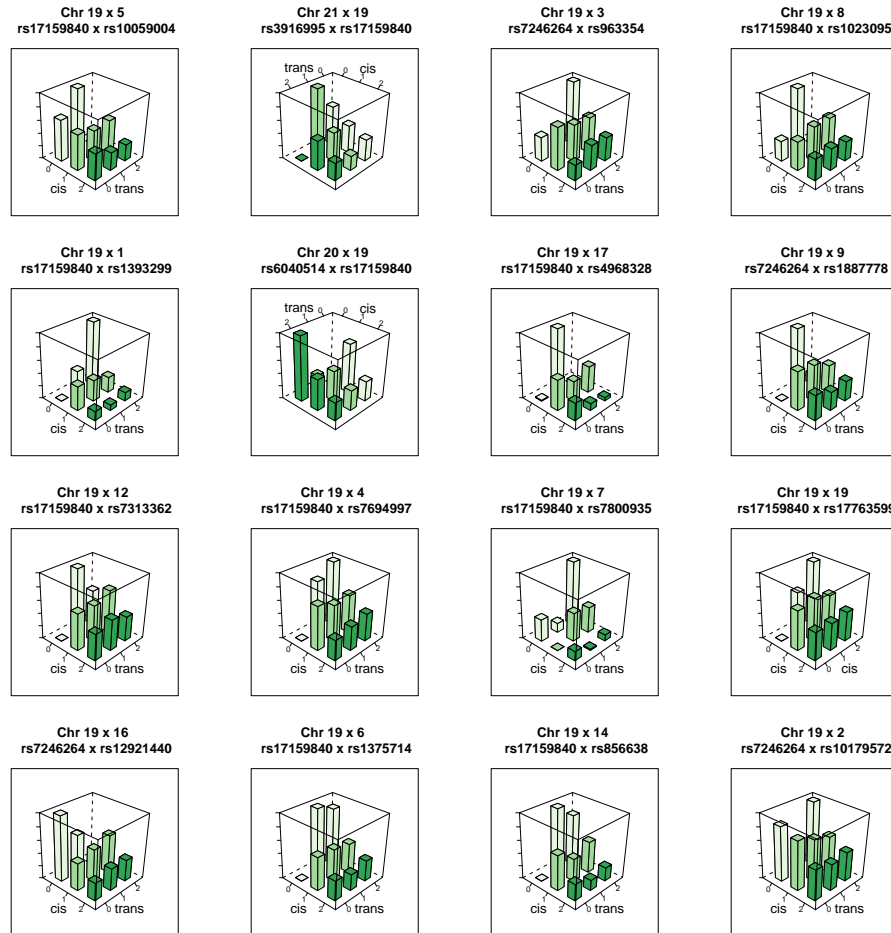


Figure S8: **Genotype-phenotype maps for 16 interactions influencing the expression of TRAPPC5** Each bar represents the mean phenotypic value for individuals in that genotype class.

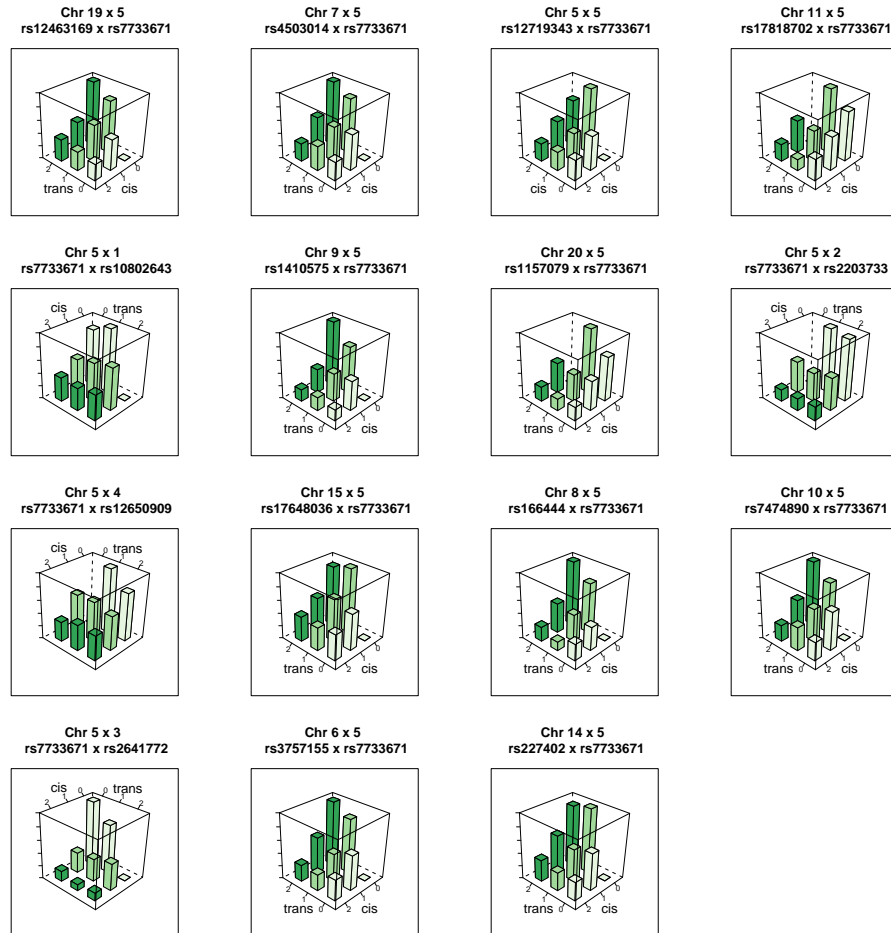


Figure S9: **Genotype-phenotype maps for 15 interactions influencing the expression of CAST** Each bar represents the mean phenotypic value for individuals in that genotype class.

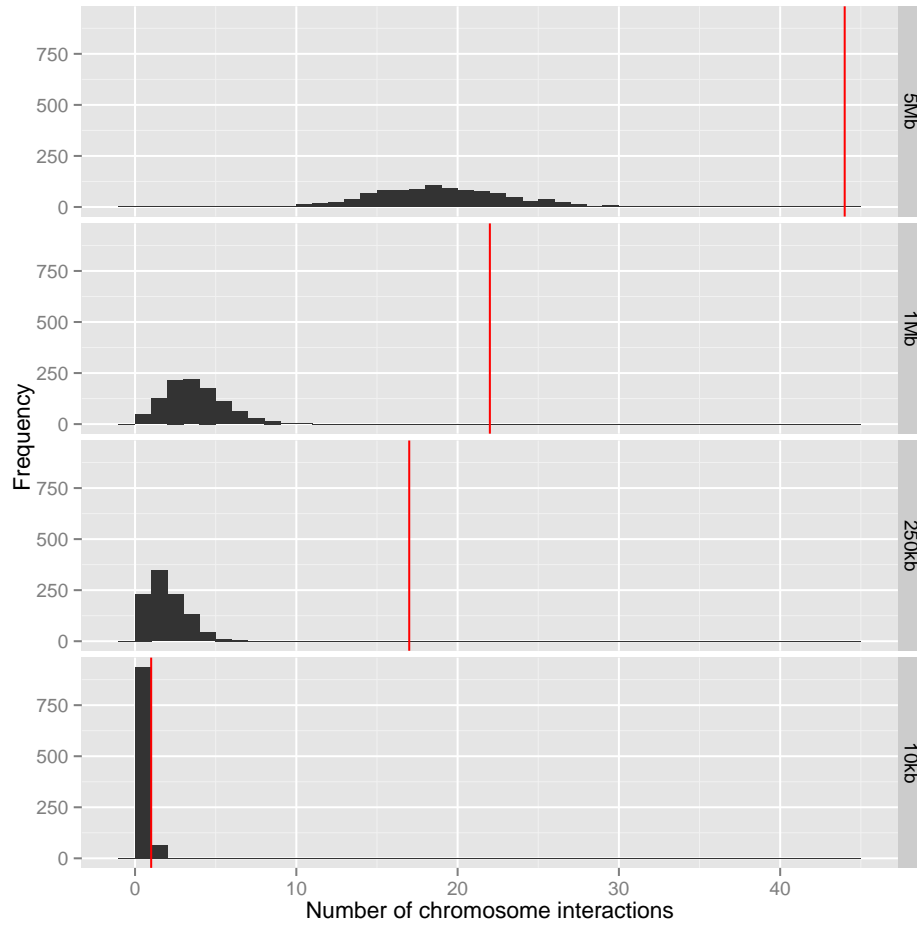


Figure S10: Number of overlaps between chromosome interactions and epistatic interactions Interacting chromosome regions may be a possible mechanism underlying epistatic interactions. The number of epistatic interactions within 20kb, 500kb, 2Mb and 10Mb of known chromosome interacting regions are shown by red vertical lines. The histograms represent the null distribution based on random sampling of 10,000 datasets for each window size.

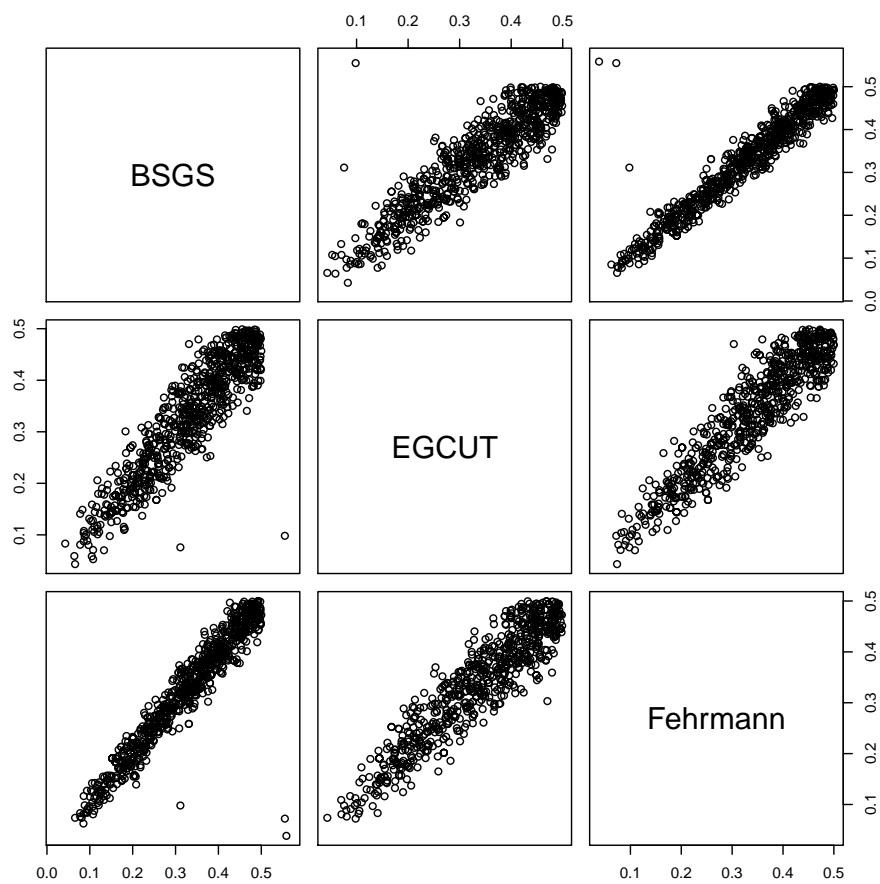


Figure S11: **Comparison of allele frequencies for 781 SNPs involved in genetic interactions across independent populations** Outliers were removed from the analysis as part of the filtering stage during replication.

5 Supplementary Tables

Table S1 – continued from previous page

Gene ID ^a	Expression trait ^b	SNP 1			SNP 2			Interaction statistic / -log ₁₀ p-values		
		rs ID	Chr.	Pos/Mb ^c	Association ^d	rs ID	Chr.	Pos/Mb ^c	Association ^d	Metag ^e
CBORF69	ILMN_1653205	rs8051751	16	7188323		rs2890452	8	86102223	CBORF59	5.79
CBORF72	ILMN_1741881	rs10122902	9	27556780	C9ORF72	rs24202910	1	24202910	C9ORF72	0.18
CAC1	ILMN_1731064	rs12765847	10	4353908		rs3738725	1	221714210	CAC1	0.96
CARD9	ILMN_1712532	rs4260763	9	139289825	INPP5E	rs654040	1	82128600		0.01
CARD9	ILMN_1712532	rs4573661	11	6026661		rs4077515	9	139266486	INPP5E	0.86
CAST	ILMN_1717234	rs112463169	20	6778978		rs7733671	5	96000269	CAST	0.42
CAST	ILMN_1717234	rs12463169	19	17321669		rs7733671	5	96000269	CAST	0.96
CAST	ILMN_1717234	rs12599264	16	81840122		rs7733671	5	96000269	CAST	2.85
CAST	ILMN_1717234	rs12719343	5	125369113		rs7733671	5	96000269	CAST	
CAST	ILMN_1717234	rs1410575	9	78255630		rs7733671	5	96000269	CAST	1.20
CAST	ILMN_1717234	rs160444	8	27392770		rs7733671	5	96000269	CAST	0.36
CAST	ILMN_1717234	rs17048036	15	27311111		rs7733671	5	96000269	CAST	1.34
CAST	ILMN_1717234	rs17818702	11	86107920		rs7733671	5	96000269	CAST	0.37
CAST	ILMN_1717234	rs227402	14	70496867		rs7733671	5	96000269	CAST	0.41
CAST	ILMN_1717234	rs2822124	21	13166804		rs7733671	5	96000269	CAST	1.09
CAST	ILMN_1717234	rs3757155	6	136458593		rs7733671	5	96000269	CAST	0.11
CAST	ILMN_1717234	rs4003014	7	31149140		rs7733671	5	96000269	CAST	
CAST	ILMN_1717234	rs4747890	10	39590078		rs7733671	5	96000269	CAST	0.07
CAST	ILMN_1717234	rs7733671	5	96000269	CAST	rs10802643	1	238120177		0.33
CAST	ILMN_1717234	rs7733671	5	96000269	CAST	rs12630909	1	170126890		1.36
CAST	ILMN_1717234	rs7733671	5	96000269	CAST	rs2203753	2	224095101		0.49
CAST	ILMN_1717234	rs7733671	5	96000269	CAST	rs2641772	3	195531841		0.78
CAST	ILMN_1717234	rs8723203	18	69173886		rs41152695	11	34115386	CAT	0.34
CDCC8B	ILMN_1692705	rs2352303	19	17093980		rs41152695	11	34115386	CAT	0.22
CDCC8B	ILMN_1722208	rs694739	17	6097235	CCDC8B	rs13771549	10	6415142	CCDC8B	0.37
CDCC8B	ILMN_1784663	rs5211834	7	80283117		rs1254900	1	8498183	YAMP8	0.11
CDCC8B	ILMN_1800940	rs7508015	11	76053374		rs1254900	1	8498183	YAMP8	0.18
CDCC8B	ILMN_1704730	rs1884655	20	23074375	CD93	rs670105	7	20755354	CD55	0.01
CDCC8B	ILMN_1704730	rs1884655	20	23074375	CD93	rs4607740	4	15782632		0.03
CDCC8B	ILMN_1704730	rs1884655	20	23074375	CD93	rs7623520	3	196721395		1.34
CDCC8B	ILMN_1704730	rs1884655	20	23074375	CD93	rs8388750	12	125145394		0.24
CDCC8B	ILMN_1704730	rs1884655	20	23074375	CD93	rs8576388	13	38434372		0.82
CDCC8B	ILMN_1704730	rs2889504	20	37771578	CD93	rs186858	20	23074375		0.27
CDCC8B	ILMN_1704730	rs4813479	20	23076914	CD93	rs10925247	1	238890903		1.67
CDCC8B	ILMN_1704730	rs4813479	20	23076914	CD93	rs2873430	8	138500554		0.22
CDCC8B	ILMN_1704730	rs4813479	20	23076914	CD93	rs4295531	18	77264432		0.51
CDCC8B	ILMN_1704730	rs4813479	20	23076914	CD93	rs7294744	17	77264432		0.11
CDCC8B	ILMN_2309796	rs901544	14	104162263		rs11655031	13	115080928	CDK16	0.14
CDCC8B	ILMN_1730928	rs200690	17	46614102	HOXB2	rs2421050	5	158043044	CDK5R1	0.95
CEACAM21	ILMN_1745949	rs4803481	19	42068556		rs4803481	19	42068556	CEACAM21	0.15
CEACAM21	ILMN_1707554	rs6505780	18	13069792	CEACAM21	rs13132719	3	134247706		0.12
CEACAM21	ILMN_1787808	rs8192935	14	101350298	CEP192	rs13079012	3	235248562	ANAPC13	0.24
CEACAM21	ILMN_2359945	rs8192935	16	55861794	CES1	rs2695290	12	102087804		0.09
CEACAM21	ILMN_2209940	rs501967	13	38838122		rs867578	12	102087804	CHPT1	0.72
CEACAM21	ILMN_2209940	rs501967	13	38838122		rs7313235	12	10132283	CLEC12A	0.44
CEACAM21	ILMN_1663142	rs429790	12	84471642		rs3903088	10	134236688	CLEC12A	0.16
CEACAM21	ILMN_2403228	rs7405054	12	10156646		rs6863172	5	173595960		0.24
CEACAM21	ILMN_1674609	rs17129799	11	96929337		rs169130	16	63121080	CLTB	0.27
CEACAM21	ILMN_1770290	rs3752237	19	1047161	ABCA7	rs169130	16	63121080	CLTB	0.07
CEACAM21	ILMN_1770290	rs3752237	19	1047161	ABCA7	rs169130	16	63121080	CLTB	0.02
CEACAM21	ILMN_1654545	rs4333645	8	145569535		rs1455268	4	61738094		1.39
CEACAM21	ILMN_1682928	rs12596791	16	26115562		rs2455884	7	29188475	CPVL	0.01
CEACAM21	ILMN_1682928	rs12596791	16	26115562		rs2455884	7	29188475	CPVL	0.57

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Table S1 – continued from previous page

Gene ID ^a	Expression trait		SNP 1		SNP 2		Interaction statistic / -log ₁₀ p-values				Distance / Mb ^b				
	Probe ID ^b	Chr.	rs ID	Chr.	Pos/Mb ^c	Association ^d	rs ID	Chr.	Pos/Mb ^c	Association ^d		BSGS ^e	Fehrmann ^f	EGCUT ^g	Meta ^g
CPVL	ILMN-1682928	7	rs2835998	21	39202070		rs245884	7	29185475	CPVL	5.55	0.19	0.03	0.04	0.033
	ILMN-1813256	2	rs2131290	4	188539908	CRLS1	rs1531133	7	46843651	CRPT	5.47	0.28	0.10	0.12	
CRUS1	ILMN-1737685	20	rs6139887	20	5986234		rs1473927	5	62406408		6.18	0.10	0.36	0.15	0.033
	ILMN-1761797	21	rs9979356	21	43230974		rs3761385	21	45196355		11.99	25.20	16.72	42.27	
CTNNA1	ILMN-1804854	5	rs924943	18	69000505		rs176382	5	138226707	CTNNA1	5.74	0.02	0.41	0.11	0.040
	ILMN-1696347	11	rs2457684	11	88139983	CTSC	rs7079264	10	10679892		5.67	0.92	0.74	0.03	
CTSC	ILMN-1696347	11	rs7532236	22	26250645		rs1728352	11	88087357	CTSC	5.84	0.49	0.73	0.03	0.040
	ILMN-2242463	11	rs7930237	11	88117962		rs12784396	10	102027407		7.16	18.76	15.06	33.53	
CWFI9L1	ILMN-1651886	10	rs7108734	11	11450027		rs12784396	10	102027407	CWFI9L1	5.42	0.21	0.01	0.03	0.040
	ILMN-1712305	4	rs2592948	4	129994690		rs888427	2	172366120	CYBRD1	5.89	0.23	0.53	0.34	
CYBRD1	ILMN-1712305	2	rs7852475	9	140698856		rs888427	2	172366120	CYBRD1	5.68	0.20	0.02	0.04	0.040
	ILMN-2087692	2	rs11257679	10	12318284		rs888427	2	172366120	CYBRD1	5.81	0.39	1.87	1.47	
CYBRD1	ILMN-2087692	2	rs6137908	20	23344590		rs888427	2	172366120	CYBRD1	5.53	0.05	0.83	0.36	0.040
	ILMN-2087692	2	rs888427	2	172366120	CYBRD1	rs7591849	2	160112881		5.85	0.87	0.10	0.44	
CYP27A1	ILMN-1704985	2	rs6021982	20	36571928		rs933994	2	219650616	CYP27A1	5.42	0.29	0.86	0.60	12.255
	ILMN-2128428	5	rs7778910	17	110451383		rs835223	5	39581357	DAB2	5.44	0.48	0.41	0.44	
DAB2	ILMN-1811648	17	rs9900173	17	133111688		rs1343244	9	82476988		5.12	0.00	0.58	0.14	0.040
	ILMN-1690982	22	rs9760102	22	24248761	DDT	rs275341	3	18470298		5.62	0.64	0.23	0.42	
DDT	ILMN-1797001	9	rs4537097	11	123962645		rs7042042	7	32451144		5.31	0.61	0.29	0.44	0.040
	ILMN-1783996	1	rs10120023	9	137810259	COQ10A	rs10120023	9	137810259	COQ10A	5.37	0.08	0.41	0.16	
DEM1	ILMN-1733998	1	rs12363827	13	106703727		rs7566044	2	169960422	DHRS9	6.39	0.77	0.02	0.29	0.040
	ILMN-1733998	2	rs1511956	12	89468283		rs7566044	2	169960422	DHRS9	6.00	0.06	1.17	0.58	
DHRS9	ILMN-2384181	2	rs1328529	12	177153535		rs2161037	2	169893419	DHRS9	6.48	0.37	0.34	0.32	0.040
	ILMN-2384181	2	rs2831914	21	29959453		rs2161037	2	169893419	DHRS9	5.51	0.88	0.04	0.37	
DHRS9	ILMN-2384181	4	rs77661304	4	187776431		rs1169322	12	50610976	LASS5	7.64	0.05	0.11	0.10	0.040
	ILMN-1755589	12	rs11080134	17	29161503	LASS5	rs2872008	7	153134888	LASS5	4.87	0.30	0.58	0.19	
DIP2B	ILMN-1755589	12	rs1166935	12	50636364		rs334593	12	20310438		4.31	0.37	0.37	0.19	0.040
	ILMN-1755589	12	rs3383835	12	50536364	LASS5	rs184654	8	15214154	LASS5	5.03	0.09	0.02	0.01	
DIP2B	ILMN-1755589	12	rs3383835	12	50536364	LASS5	rs184654	8	15214154	LASS5	5.03	0.09	0.02	0.01	0.040
	ILMN-1755589	12	rs3383835	12	50536364	LASS5	rs184654	8	15214154	LASS5	5.03	0.09	0.02	0.01	
DIP2B	ILMN-1755589	12	rs3383835	12	50536364	LASS5	rs184654	8	15214154	LASS5	5.03	0.09	0.02	0.01	0.040
	ILMN-1755589	12	rs3383835	12	50536364	LASS5	rs184654	8	15214154	LASS5	5.03	0.09	0.02	0.01	
DNAB1B6	ILMN-1793770	7	rs2288842	15	177094348		rs12437378	12	15716199	DNAB1B6	5.70	0.23	1.45	0.97	0.052
	ILMN-2109708	22	rs12437378	12	15716199	ECGF1	rs3775539	7	6320360	DNAB1B6	4.17	1.58	0.27	1.12	
EGCF1	ILMN-2109708	22	rs12437378	12	15716199	ECGF1	rs3775539	7	6320360	DNAB1B6	4.17	1.58	0.27	1.12	0.052
	ILMN-1671568	1	rs4324091	22	50971266		rs1560972	3	6320360	DNAB1B6	4.17	1.58	0.27	1.12	
EGHDC2	ILMN-1671568	1	rs4324091	22	50971266		rs1560972	3	6320360	DNAB1B6	4.17	1.58	0.27	1.12	0.052
	ILMN-1720083	15	rs5092637	22	50971266		rs1560972	3	6320360	DNAB1B6	4.17	1.58	0.27	1.12	
EHD4	ILMN-1720083	15	rs5092637	22	50971266		rs1560972	3	6320360	DNAB1B6	4.17	1.58	0.27	1.12	0.052
	ILMN-1719380	14	rs6567288	18	60218334		rs11206043	1	53402552	EGHDC2	6.19	0.22	0.35	0.22	
EIF2B2	ILMN-1719380	14	rs6567288	18	60218334		rs11206043	1	53402552	EGHDC2	6.19	0.22	0.35	0.22	0.052
	ILMN-1704522	17	rs7216490	17	7221707	EIF5A	rs1043166	15	42192040	EIF2B2	6.98	0.90	0.47	0.79	
EIF5A	ILMN-1794522	17	rs7216490	17	7221707	EIF5A	rs1269096	14	99603119		5.44	0.56	0.08	0.24	0.052
	ILMN-1745522	17	rs7216490	17	7221707	EIF5A	rs1553474	2	49359676		5.55	0.28	0.59	0.41	
EIF5A	ILMN-1794522	17	rs7216490	17	7221707	EIF5A	rs1269096	14	99603119		5.55	0.28	0.59	0.41	0.052
	ILMN-1794522	17	rs7216490	17	7221707	EIF5A	rs1269096	14	99603119		5.55	0.28	0.59	0.41	
EIF5A	ILMN-1794522	17	rs7216490	17	7221707	EIF5A	rs1269096	14	99603119		5.55	0.28	0.59	0.41	0.052
	ILMN-1794522	17	rs7216490	17	7221707	EIF5A	rs1269096	14	99603119		5.55	0.28	0.59	0.41	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	0.052
	ILMN-2353633	19	rs2827076	21	23196249		rs4471434	11	126387391	EMR2	5.52	0.05	1.12	0.52	
EMR2	ILMN-2353633	19	rs2827076	21	23196249		rs4471434								

Table S1 – continued from previous page

Gene ID ^a	Probe ID ^b	Chr.	rs ID	Chr.	SNP 1	Pos/Mb ^c	Association ^d	rs ID	Chr.	SNP 2	Pos/Mb ^c	Association ^d	BSGS ^e	Interaction statistic ^f	EGCUT ^g	− log ₁₀ <i>p</i> -value ^h	Distance / Mb ⁱ
FE2Z	ILMN-1739586	2	rs2356400	19	44321776	19		rs13406184	2	36791226	FE2Z	FE2Z	5.78	0.14	0.33	0.16	
FE2Z	ILMN-1739586	2	rs969010	4	169963132	4		rs11691600	6	36810133	FE2Z	FE2Z	6.59	0.14	0.28	0.14	
FGD2	ILMN-2115005	6	rs4803848	19	46203050	19		rs831486	6	37001267	FGD2	FGD2	5.69	0.12	0.25	0.11	
FGD2	ILMN-2115005	6	rs902634	12	133943951	12		rs351498	6	36999652	FGD2	FGD2	5.49	1.20	0.11	0.66	68.867
FL20489	ILMN-1761073	12	rs17036706	12	177036706	12	FL20489	rs831486	6	41699526	FL20489	FL20489	5.81	0.06	0.70	0.29	
FL20489	ILMN-1761073	12	rs17036706	12	177036706	12		rs3782908	12	48169526	FL20489	FL20489	5.79	0.19	0.13	0.04	
FL20489	ILMN-1778144	12	rs7021190	15	70929121	15		rs3782908	12	48169526	FL20489	FL20489	5.79	0.19	0.13	0.04	
FL20489	ILMN-1778144	12	rs484440	12	97033126	12		rs3782908	12	48169526	FL20489	FL20489	6.04	0.31	0.47	0.36	
FL20489	ILMN-1778144	12	rs7204135	16	50626195	12		rs3782908	12	48169526	FL20489	FL20489	6.90	0.38	0.17	0.21	
FL20489	ILMN-1763663	16	rs9325634	21	43818790	16		rs3782908	12	50116594	FL20489	FL20489	6.04	0.14	0.95	0.53	
FL20489	ILMN-1763663	16	rs71127627	14	107276627	16		rs6906101	6	36676710	FL20489	FL20489	5.48	0.39	0.06	0.13	3.962
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6	FL20489	rs6906101	6	36676710	FL20489	FL20489	5.48	0.39	0.06	0.13	0.063
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	
FL20489	ILMN-2123450	6	rs6906101	14	107276627	6		rs9892064	17	808297903	FL20489	FL20489	16.16	28.24	29.39	59.95	

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Gene ID ^a	Expression trait		SNP 1			SNP 2			Interaction statistic ^c		Distance / Mb	
	rs ID	Chr.	Pos/Mb/c	Association ^d	rs ID	Chr.	Pos/Mb/c	Association ^d	BSGS ^e	Fehrmann ^f		
Probe ID ^b	rs43925535	4	41513423		rs13069559	3	152187431	MBNL1	8.39	0.02	4.33	3.02
	rs47358380	8	895841		rs13069559	3	152187431	MBNL1	6.74	0.32	4.21	3.38
	rs49393558	18	46278591		rs1522374	3	152235530		7.72	0.03	0.27	0.07
	rs6128397	3	57253132		rs16864367	3	152234166		7.22	1.34	1.15	1.73
	rs7701738	5	22101322		rs13069559	3	152187431	MBNL1	7.92	2.55	7.89	9.28
	rs6079849	20	15462611		rs2051344	18	74715653	MBP	6.26	0.10	0.03	0.02
	rs139568	22	42210985		rs2051344	18	74715653	MBP	5.56	0.03	0.23	0.05
	rs2051344	18	74715653	MBP	rs1125539	3	155204939		5.79	0.02	0.76	0.27
	rs2051344	18	74715653	MBP	rs2619046	6	55097534		6.03	0.15	0.50	0.26
	rs4805021	19	33436367		rs2051344	18	74715653	MBP	5.82	0.03	0.47	0.14
Gene ID ^a	rs8092433	18	74747424		rs49890876	18	74732087	MEGF9	4.63	7.06	21.91	28.73
	rs13039689	20	51922071		rs4846085	1	12050634	MFN2	5.76	0.61	0.25	0.41
	rs7989895	13	109401737	MGC13057	rs12718598	7	50428445	MGC13057	5.81	0.13	0.30	0.14
	rs674608	18	82607772		rs12718598	7	50428445	MGC13057	5.07	0.11	1.03	0.50
	rs8058318	16	69628245		rs2660665	8	137526799		4.17	0.05	0.08	0.02
	rs845787	20	51977931		rs4147592	5	165600146	MGST3	5.45	0.57	0.27	0.40
	rs740441	17	25779644	MPZL2	rs11771552	7	154708716		5.90	0.01	0.23	0.04
	rs1805	11	118076069		rs1805	11	118076069	MPZL2	5.64	0.97	1.08	1.35
	rs7316716	12	19953193		rs750495	5	1782046	MRPL36	6.89	0.34	0.18	0.19
	rs17469061	16	80641040		rs2636095	10	102740503	MRPL36	5.71	0.26	0.14	0.22
Gene ID ^a	rs6564769	16	80641040		rs3811188	14		MRPL52	6.56	0.14	0.44	0.64
	rs1950857	14	267102371		rs722269	6	42194916	MRPS10	7.48	0.46	0.70	0.22
	rs10955512	8	110202230		rs2395803	6	42158596	MRPS10	6.85	0.31	0.63	0.46
	rs11698155	20	5063214		rs132717993	6	42164401	MRPS10	6.21	0.41	0.25	0.28
	rs1420537	16	52453567	MTNR10	rs12431444	14	42068689		5.18	1.87	2.86	3.61
	rs1738375	15	31215935		rs11160227	14	95514596		6.31	0.46	0.52	0.50
	rs459498	21	42795027		rs4973801	21	26706382		5.83	0.11	0.50	0.23
	rs459498	21	42795027		rs130120	21	29363604	MYBPC3	6.78	0.29	0.92	0.65
	rs10134030	14	61593110		rs1317149	11	47486885	MYBPC3	5.56	0.13	0.46	0.23
	rs7322708	13	109530661	MYOM1	rs1734423	11	47325947	MYBPC3	5.70	0.04	0.08	0.02
Gene ID ^a	rs1958073	18	8247256		rs2734722	8	134483237	NABP1	5.92	0.74	1.15	1.40
	rs1958073	18	8247256		rs2734722	8	134483237	NABP1	5.92	0.74	1.15	1.40
	rs2707574	7	147638723		rs2707574	7	147638723	NAAA	5.45	0		

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Table S1 – continued from previous page

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Table S1 – continued from previous page

Gene ID ^a	Expression trait		SNP 1		Pos/Mb ^c	Association ^d	SNP 2		Pos/Mb ^c	Association ^d	Interaction statistic / -log ₁₀ p-values				Distance / Mb ^h
	Probe ID ^b	Chr.	rs ID	Chr.	Pos/Mb ^c	Association ^d	rs ID	Chr.	Pos/Mb ^c	Association ^d	BSGS ^e	Fehrmann ^f	EGCUT ^g	Meta ^g	
REBE	ILMN-1802380	1	rs4982958	14	24987865		rs301819	1	8501786	REBE	5.66	0.61	1.23	1.17	
REBE	ILMN-1802380	1	rs7697290	4	135248366		rs301819	1	8501786	REBE	5.74	0.14	0.10	0.06	
REBE	ILMN-2327795	1	rs11085829	19	13174312		rs301819	1	8501786	REBE	5.12	0.21	0.33	0.21	
REBE	ILMN-2327795	1	rs3852011	3	112844086	RNASE6	rs301819	1	8501786	REBE	5.71	0.08	0.60	0.26	
RNASE6	ILMN-1780533	14	rs11628398	14	8106521		rs7324365	13	100601327	RNASE6	5.48	0.42	0.21	0.26	
RNASE6	ILMN-1780533	14	rs6603134	19	8106521		rs11628398	14	21182800	RNASE6	5.11	0.09	0.22	0.08	
RNF167	ILMN-1794726	17	rs2382330	17	4875566		rs4848487	13	54668512		4.37				
RNF167	ILMN-1794726	17	rs400668	17	4839930		rs11706900	3	36348968		5.59	0.71	0.46	0.64	
RNFEP	ILMN-1738347	1	rs1107121	17	46127549		rs2819365	1	201983242		6.27	0.11	0.30	0.13	
RNFEP	ILMN-1738347	1	rs8071611	17	67153386		rs2819365	1	201983242		4.32	1.48	0.52	1.28	
RPL13	ILMN-2413278	16	rs352935	16	89648580		rs2965817	16	89513234		4.98	3.79	14.41	17.24	
RPL23AP7	ILMN-2222750	2	rs1401202	16	80320056		rs4849261	2	114450028	RPL23AP7	5.55	0.13	0.73	0.38	
RPL36AL	ILMN-2186933	14	rs3007033	14	50103816		rs17495030	9	138035083		5.46	0.09	0.06	0.02	
RPL36AL	ILMN-2186936	14	rs4900928	14	50020817	RPL36AL	rs1502991	6	66137260		5.86	0.32	0.20	0.19	
RPL8	ILMN-1764721	8	rs2958482	8	145984615	RPL8	rs1619856	1	234585790		4.59	0.10	0.37	0.15	
RPL8	ILMN-1764721	8	rs4143674	20	4741304		rs2958482	8	145984615	RPL8	4.33	0.13	0.45	0.22	
SEC13	ILMN-3297880	3	rs4889214	16	80913946		rs696221	3	10342876	SEC13	6.48	0.22	1.73	1.17	
SEMA4A	ILMN-1702787	1	rs17085428	5	95388015		rs7695	1	156147326	SEMA4A	5.70	0.02	0.51	0.15	
SEMA3	ILMN-1694027	11	rs12147460	14	104412137		rs684856	11	94906111	SEMA3	5.50	0.31	0.06	0.10	
SEMA3	ILMN-1694027	11	rs355391	15	46391793	SEMA3	rs684856	11	94906111	SEMA3	5.67	0.31	0.06	0.10	
SEMA3	ILMN-1694027	11	rs684856	11	46391793		rs7004947	8	134606425	SEMA3	5.60	0.21	0.51	0.31	
SH3BGL2	ILMN-1762764	6	rs10838191	11	43893658		rs1354034	3	56849749	PPBP	5.52	0.70	0.12	0.35	
SH3BGL2	ILMN-1762764	6	rs2545385	5	66383979		rs1354034	3	56849749	PPBP	5.97	0.20	0.51	0.30	
SH3BGL2	ILMN-1762764	6	rs6845304	4	88280592		rs1354034	3	56849749	PPBP	5.23	0.32	0.71	0.53	
SH3BGL2	ILMN-2158336	9	rs1034210	21	18196922		rs17455517	9	131785369	SH3BGL2	7.40	0.22	0.18	0.13	
SIRPG	ILMN-1771801	20	rs1535883	20	1612819	SIRPG	rs6842739	14	60489510		5.74	0.29	0.18	0.17	
SLC22A18	ILMN-2382505	11	rs11673260	19	52181798		rs367035	11	2923826	SLC22A18	5.47	0.09	0.24	0.09	
SLC22A18	ILMN-2382505	11	rs367035	19	52181798	SLC22A18	rs3110874	7	153224179		5.70	0.15	0.10	0.06	
SLC22A18	ILMN-2382505	11	rs367035	11	2923826		rs3772054	2	241678528	SLC22A18	6.15	0.39	0.13	0.19	
SLC41A3	ILMN-2356111	3	rs1912136	11	24616743		rs6771703	3	125801067	SLC41A3	5.88	1.10	0.82	1.24	
SLC45A4	ILMN-1747778	8	rs6985508	8	142337734	SLC45A4	rs7701916	5	174598073	SLC45A4	5.95	0.86	0.07	0.40	
SLC46A3	ILMN-1658639	13	rs949805	17	55602091		rs7981190	13	29259349	SLC46A3	5.52	0.09	0.58	0.26	
SMG7	ILMN-1706553	1	rs8035259	15	97403923		rs10911353	1	183489203	SMG7	6.52	0.17	0.09	0.06	
SMOX	ILMN-1753380	20	rs118315	20	4161500	SMOX	rs11677815	2	65800982		5.68	0.39	0.62	0.52	
SNHG8	ILMN-3309349	4	rs1105321	9	133050233		rs705837	4	119225940	SNHG8	6.11				
SNORD14A	ILMN-1799381	11	rs1520429	15	46259108		rs214097	11	17291499	SNORD14A	6.60	0.29	1.03	0.72	
SNORD14A	ILMN-1799381	11	rs2634462	11	17339127		rs6486334	11	17015557	SNORD14A	7.31	13.11	10.96	23.22	
SNORD89	ILMN-3238662	2	rs10445863	2	115929241		rs750783	2	101889306	SNORD89	6.08				
SNORD89	ILMN-3238662	2	rs11605822	11	122986326		rs750783	2	101889306	SNORD89	5.96				
SNORD89	ILMN-3238662	2	rs2135064	5	26778066		rs750783	2	101889306	SNORD89	6.33				
SNUPN	ILMN-1733932	15	rs1346466	21	46376528	SNUPN	rs17185362	16	81888905		6.45				
SNUPN	ILMN-2364535	15	rs1346466	21	46376528		rs1472075	3	193706323		5.59	0.34	0.00	0.06	
SPATA5L1	ILMN-1729179	15	rs1313620	19	41117869		rs1472075	3	193706323	SPATA5L1	5.44	0.67	0.12	0.33	
STARD10	ILMN-2210752	7	rs4073164	14	104947517		rs1006620	11	72509713	STARD10	5.85	0.34	0.00	0.06	
STYXL1	ILMN-2345142	20	rs2221406	13	90174526		rs17685	7	75616105	STYXL1	5.51	0.17	0.31	0.57	
SULT1A4	ILMN-2336133	16	rs1463965	18	74332954	SULF2	rs392994	4	180439236	SULT1A4	7.05	0.46	0.24	0.30	
SULT1A4	ILMN-2336133	16	rs2436657	20	40119768		rs3785354	16	28550667	TUFM	5.83	0.01	0.05	0.00	
SURF6	ILMN-1778032	9	rs6099626	21	56013994		rs3785354	16	28550667	TUFM	7.05	0.01	0.05	0.00	
SYTL2	ILMN-2336609	11	rs1375719	13	103410782		rs3118663	9	136281753	SURF6	6.14	0.26	0.16	0.14	
THBS3	ILMN-1804663	1	rs1939875	11	95422867		rs485485	11	85495269	SYTL2	5.47	0.28	0.31	0.24	
THBS3	ILMN-1804663	1	rs1939875	11	95422867		rs4072037	1	155162067	THBS3	5.55	0.03	0.15	0.03	
TIPRL	ILMN-1781457	1	rs8014956	14	20687978		rs20449805	1	155194980	TIPRL	5.65	0.31	0.76	0.55	
TIPRL	ILMN-1781457	1	rs2823245	21	16745523		rs1320993	1	168154599	TIPRL	5.22	0.07	0.40	0.15	

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Table S1 – continued from previous page

Gene ID ^a		Expression trait		SNP 1		SNP 2		Interaction statistic / -log ₁₀ p-values		Meta ^g	Distance / Mb ^h			
TMED4	Probe ID ^b	Chr.	rs ID	Chr.	Pos / Mb ^c	Association ^d	rs ID	Chr.	Pos / Mb ^c			Association ^d	BSGS ^e	Fehrmann ^f
TMED4	ILMN-1804148	7	rs19340400	11	132389627		rs17725246	7	44581986	TMED4	3.70	0.06	1.34	0.70
TMEM149	ILMN-1786426	19	rs28390113	21	47248981		rs8106959	19	36219525	TMEM149	8.11	0.16	0.48	0.26
TMEM149	ILMN-1786426	19	rs5762235	22	27925288		rs8106959	19	36219525	TMEM149	6.79			
TMEM149	ILMN-1786426	19	rs6090518	20	43207005		rs8106959	19	36219525	TMEM149	11.09	0.76		
TMEM149	ILMN-1786426	19	rs807491	19	36268923	SNX26	rs7254601	19	36147315	TMEM149	12.16	81.55	45.78	145.78
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs10508289	10	47991159		8.12	1.55	3.09	3.07
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs10819626	9	133025736		8.02	0.40	0.99	0.80
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs10937361	3	188350436		5.39	3.61	1.18	3.78
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs1401098	12	128884559		7.37	2.41	1.00	2.52
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs1557335	18	64268976		6.95	0.08	0.07	0.03
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs17719594	14	90932398		6.93	3.06	0.77	2.87
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs1843357	8	13822381		6.21	3.72	3.33	6.00
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs2351458	4	113317583		7.30	0.04	9.61	8.00
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs2539000	7	147619772		6.70	1.57	1.52	2.27
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs2731711	5	171792273		5.92	0.19	0.33	0.19
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs4711728	11	129595460		8.89	0.90	3.62	3.51
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs6718480	2	233879066		8.55	3.31	5.15	7.36
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs6926382	2	161683974		5.80	3.06	8.80	10.72
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs7213338	17	80357420		5.49	0.07	3.14	2.10
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs914940	1	242889492		6.22	3.36	6.96	9.20
TMEM149	ILMN-1786426	19	rs8106959	19	36219525	TMEM149	rs9509428	13	21473952		9.44	0.10	5.75	4.47
TMEM63A	ILMN-1719649	1	rs1254086	13	72890603		rs4149226	1	226027323	TMEM63A	5.60			
TMEM80	ILMN-1768482	11	rs1548475	9	58058246		rs4963126	11	65845	TMEM80	5.79	0.64	0.12	0.32
TMPO3	ILMN-1683811	7	rs1537146	9	4859303		rs10488630	7	128593948	IRF5	5.61	0.11	0.15	0.07
TMPO3	ILMN-1683811	7	rs199793	20	22287303		rs10488630	7	128593948	IRF5	5.52	1.03	0.17	0.62
TRAF2	ILMN-1731043	7	rs7776572	7	23528927		rs11770192	7	23498358		8.23	3.19	1.89	4.09
TRAPPC4	ILMN-1814650	11	rs1278760	13	113531675		rs3916581	11	11887887	TRAPPC4	5.61	0.28	0.40	0.29
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs10059004	5	166970604	TRAPPC4	5.52	0.93	0.01	0.36
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs1023095	8	132022957		6.92	0.37	1.60	1.07
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs1375714	6	156404902		7.79	0.12	0.18	0.07
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs1393299	1	242329791		6.43	0.63	0.47	0.59
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs17763599	19	2369415		6.38	0.21	0.24	0.16
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs4968328	17	57495457		6.51	0.50	0.38	0.44
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs7313362	12	129644342		7.08	0.04	0.65	0.25
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs7694997	4	9947811		5.86	0.20	0.36	0.22
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs7800935	7	146690926		6.27	0.15	0.33	0.16
TRAPPC5	ILMN-2372639	19	rs17159840	19	7758194	TRAPPC5	rs856638	14	85439550		6.73	0.24	0.07	0.08
TRAPPC5	ILMN-2372639	19	rs30708	22	22740855		rs17159840	19	7758194	TRAPPC5	7.58			
TRAPPC5	ILMN-2372639	19	rs3916995	19	45128451		rs17159840	19	7758194	TRAPPC5	7.73	0.85	0.78	1.01
TRAPPC5	ILMN-2372639	19	rs6040514	20	11272861		rs17159840	19	7758194	TRAPPC5	8.10	0.51	0.55	0.56
TRAPPC5	ILMN-2372639	19	rs7246264	19	7762978		rs10179572	2	228504503		6.71	0.14	0.02	0.02
TRAPPC5	ILMN-2372639	19	rs7246264	19	7762978		rs12921440	16	30408795		7.34	0.14	0.26	0.13
TRAPPC5	ILMN-2372639	19	rs7246264	19	7762978		rs1887778	3	134635088	RAPGEF1	7.05	0.08	0.86	0.40
TRAPPC5	ILMN-2372639	19	rs7246264	19	7762978		rs963354	3	157393770		7.41	0.36	0.90	0.69
TREM1	ILMN-1688231	6	rs10862975	12	85749398		rs2395771	6	41264577	TREM1	5.42	0.11	0.25	0.11
TREM1	ILMN-1688231	6	rs2527180	17	158808416		rs2395771	6	41264577	TREM1	5.92	1.20	1.23	1.69
TRIM38	ILMN-1697971	6	rs2527180	17	158808416		rs2032447	6	26044369	TRIM38	6.46	0.04	0.91	0.39
TSPAN14	ILMN-1785060	10	rs968726	7	27194634	MYBPC3	rs10748526	10	82273079	TSPAN14	6.00	0.07	0.18	0.06
TSPAN32	ILMN-1718621	11	rs10838738	11	47663049	MYBPC3	rs12800098	11	2317951	TSPAN32	5.01			
TSPAN32	ILMN-2389070	11	rs12800098	11	2317951	TSPAN32	rs620607	6	137947208	TSPAN32	5.51			
TRIP	ILMN-323126	22	rs140522	22	50971266	ECGF1	rs1198819	2	238746880	TRIP	6.34			
TRIP	ILMN-323126	22	rs470119	22	50966914	ECGF1	rs4783126	16	85147633	TRIP	6.13			

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Expression trait			SNP 1			SNP 2			Interaction statistic / -log ₁₀ p-values			Distance / Mb			
Gene ID ^a	Probe ID ^b	Chr.	rs ID	Chr.	Pos /Mb ^c	Association ^d	rs ID	Chr.	SNP 2	Association ^d	BSGS ^e	Fehrmann ^f	EGCUT ^g	Meta ^g	Distance / Mb ^h
UBASH3A	ILMN-2338348	21	rs1893592	21	43855067	UBASH3A	rs7201194	16	83600397	5.91	0.59	0.42	0.52		
UBASH3A	ILMN-2338348	21	rs1893592	21	43855067	UBASH3A	rs7512594	16	214514361	6.01	0.48	1.29	1.10		
USP36	ILMN-1697227	17	rs2279308	17	76794981	USP36	rs7225546	17	75151717	5.71	0.03	0.14	0.03		1.643
VASP	ILMN-1743646	19	rs1264226	19	40063167		rs2276470	19	45974668	5.09	0.94	5.14	4.95		0.088
VNN2	ILMN-1678939	6	rs10435352	7	103252718		rs1883613	6	133077063	VNN2		5.64	0.84	0.15	0.46
VNN2	ILMN-1678939	6	rs13044386	20	9116155		rs1883617	6	133072650	VNN2		5.44	0.39	0.69	0.57
VNN2	ILMN-1678939	6	rs134447	22	49927332		rs1883617	6	133072650	VNN2		5.72			
VNN2	ILMN-1678939	6	rs216495	11	16834510		rs1883617	6	133072650	VNN2		5.77	0.33	0.19	0.19
VNN3	ILMN-1804935	6	rs10278073	7	151662184		rs2267932	6	133067782	VNN3		6.44	0.16	0.74	0.41
VNN3	ILMN-1804935	6	rs1443946	8	73006453		rs2267932	6	133067782	VNN3		5.74	0.23	0.48	0.31
VNN3	ILMN-1804935	6	rs348462	9	73547169		rs2267952	6	133067782	VNN3		6.44	0.31	0.17	0.17
VNN3	ILMN-1804935	6	rs7157055	14	83262064		rs2267952	6	133067782	VNN3		5.82	0.03	0.19	0.04
VNN3	ILMN-2387680	6	rs2823165	21	5694253		rs2267952	6	133067782	VNN3		6.12	0.73	1.15	1.21
VNN3	ILMN-2387680	6	rs9596457	13	51692548		rs2267952	6	133067782	VNN3		4.83	0.46	0.05	0.16
VSTM1	ILMN-1763455	19	rs10500316	19	54553697	VSTM1	rs4552100	18	71024750		5.60	0.53	0.54	0.57	
VSTM1	ILMN-1763455	19	rs10500316	19	54553697	VSTM1	rs7895870	10	123095249		5.71	0.48	1.17	0.26	
VSTM1	ILMN-1763455	19	rs9628570	22	30261219		rs10500316	19	54553697	VSTM1		5.88	0.81	1.38	1.47
WDR48	ILMN-1762103	3	rs1388935	4	138927822		rs6778963	3	39091812	WDR48		5.88	0.19	0.33	0.09
WDR48	ILMN-1762103	3	rs1887778	3	134635088	RAPGEF1	rs853349	3	39067925	WDR48		5.94	0.57	1.35	1.22
WDR6	ILMN-1762103	3	rs9554833	13	102624790		rs7619193	3	39044116	WDR48		5.85	0.18	0.61	0.35
WDR6	ILMN-1669484	3	rs12362253	11	123571708		rs7715581	3	49194351	WDR6		4.86	1.64	1.43	2.25
XAF1	ILMN-233053	17	rs1535031	21	9673170	XAF1	rs12591171	15	93119799		5.86	2.38	0.17	1.63	
ZFP90	ILMN-1680573	16	rs909446	17	37040648		rs182968	16	68573945	ZFP90		5.79	0.09	0.36	0.15
ZNF500	ILMN-1700238	16	rs4282723	22	48283177		rs2290560	16	4799041	ZNF500		5.29	0.67	0.27	0.46
ZYX	ILMN-1701875	7	rs6056281	20	8935312		rs2242601	7	143093824	ZYX		6.04	0.26	0.01	0.05

^a Phenotypes are expression levels of RefSeq Genes^b Illumina probe ID used to measure gene expression^c Physical SNP position in base pairs (HG19)^d RefSeq Gene ID of gene expression level that is influenced by the SNP (BSGS discovery dataset, significance threshold = 1.29 × 10⁻¹¹)^e Interaction - log₁₀ p-value from discovery dataset^f Interaction - log₁₀ p-value from replication dataset^g Interaction - log₁₀ p-value from meta analysis of replication datasets only^h Distance in Mb between interacting SNPs for *cis-cis* acting SNP pairs

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