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Supplementary appendix

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Supplementary Material for Ndila *et al.*

Human candidate gene polymorphisms and risk of severe malaria in children in Kilifi, Kenya: a case-control association study

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#	Gene Symbol	Gene Description	SNP ID	Chr	Position	Removed in QC
1	AJAP1	Adherens junctions associated protein 1	rs113788643	1	4817664	
			rs6674631	1	4834821	
2	LPHN2	Latrophilin 2	rs146428334	1	81688714	
			rs72933304	1	81726138	
			rs72933310	1	81727427	
			rs72933350	1	81751439	
			rs4650365	1	81770827	
3	GBP7	Guanylate binding protein 7	rs1803632	1	89582690	
4	DARC	Duffy blood group, atypical chemokine receptor	rs2814778	1	159174683	
			rs55868763	1	203652140	
			rs1541255	1	203652141	
5	ATP2B4	ATPase, Ca ²⁺ transporting, plasma membrane 4	rs10900585	1	203654024	
			rs4951074	1	203660781	
			rs3753036	1	203677250	
			rs3024500	1	206940831	
6	IL10	Interleukin 10	rs1800896	1	206946897	
			rs1800890	1	206949365	
			rs17047660	1	207782856	
7	CR1	Complement receptor 1	rs17047661	1	207782889	
			rs1371478	2	4901589	
8	LOC727982	Long intergenic non-protein coding RNA 1249	rs1371474	2	4909777	
			rs10188961	2	4926593	
			rs973128	2	20332487	
9	LAPTM4A	Lysosomal protein transmembrane 4 alpha	rs973128	2	20332487	
10	SDC1	Syndecan 1	rs11899121	2	20367973	
11	IL1A	Interleukin 1, alpha	rs17561	2	113537223	
12	IL1B	Interleukin 1, beta	rs1143634	2	113590390	
13	ZSWIM2	Zinc finger, SWIM-type containing 2	rs4316902	2	188007364	
			rs144778284	2	188012821	
14	IL17RE	Interleukin 17 receptor E	rs708567	3	9960070	
15	OXNAD1	Oxidoreductase NAD-binding domain containing 1	rs200704287	3	16407519	monomorphic
			rs79691057	3	16408251	monomorphic
			rs75180423	3	16408723	monomorphic
16	TLR9	Toll-like receptor 9	rs187084	3	52261031	
17	IL17RD	Interleukin 17 receptor D	rs6780995	3	57138419	
18	ARL14	ADP-ribosylation factor-like 14	rs76033371	3	160362359	
			rs75731597	3	160364808	
			rs74954675	3	160381509	
19	B3GALNT1	Beta-1,3-N-acetylgalactosaminyltransferase 1 (Globoside blood group)	rs12107243	3	160793678	
20	TLR1	Toll-like receptor 1	rs4833095	4	38799710	
21	TLR6	Toll-like receptor 6	rs5743810	4	38830350	
			rs5743809	4	38830514	
22	INPP4B	Inositol polyphosphate-4-phosphatase, type II, 105kda	rs77389579	4	143538511	
			rs13103597	4	143558581	
23	USP38	Ubiquitin specific peptidase 38	rs4266246	4	143971242	
			rs28459062	4	144039139	
24	GAB1	GRB2-associated binding protein 1	rs7663712	4	144261117	
25	GUSBP5	Glucuronidase, beta pseudogene 5	rs148111931	4	144540045	monomorphic
			rs184908374	4	144665753	
			rs149914432	4	144666678	
			rs186790584	4	144680140	
			rs184895969	4	144698528	
26	FREM3	FRAS1 related extracellular matrix 3	rs186873296	4	144702474	
			rs1801033	5	41199959	
			rs2706384	5	131826880	
28	IRF1	Interferon regulatory factor 1	rs2706384	5	131826880	
29	IL13	Interleukin 13	rs20541	5	131995964	
30	IL4	Interleukin 4	rs2243250	5	132009154	
31	GABBR1	Gamma-aminobutyric acid (GABA) B receptor, 1	rs192151845	6	29588309	

32	<i>HCG4</i>	HLA complex group 4	rs114980857	6	29772098	
33	<i>LTA</i>	Lymphotoxin alpha	rs2239704	6	31540141	
			rs909253	6	31540313	
			rs1799964	6	31542308	
			rs1800750	6	31542963	
34	<i>TNF</i>	Tumour necrosis factor	rs1800629	6	31543031	
			rs361525	6	31543101	
			rs3093662	6	31544189	
35	<i>HSPA1B</i>	Heat shock protein family A (HSP70) member 1B	rs6457452	6	31795550	
36	<i>SNORD48</i>	Small nucleolar RNA, C/D box 48	rs116288147	6	31803074	
37	<i>CTLA</i>	SLC44A4 (Solid carrier family 44, member 4)	rs2242665	6	31839309	
38	<i>IL20RA</i>	Interleukin 20 receptor, alpha	rs1555498	6	137325847	
			rs55958968	6	150942218	
			rs144224092	6	150973623	missingness
			rs79100774	6	150975934	
			rs114726617	6	150980481	
			rs2131263	6	150981102	
39	<i>PLEKHG1</i>	Pleckstrin homology domain containing, family G member 1	rs76924464	6	150982529	
			rs151293197	6	150994429	
			rs142712208	6	151026346	
			rs15116938	6	151046029	
			rs141555199	6	151048708	
40	<i>NOD1</i>	Nucleotide-binding oligomerization domain containing 1	rs2075820	7	30492237	
41	<i>CD36</i>	CD36 molecule (thrombospondin receptor)	rs3211938	7	80300449	
42	<i>TLR4</i>	Toll-like receptor 4	rs4986790	9	120475302	
			rs4986791	9	120475602	
			rs8176750	9	136131057	
			rs56390333	9	136131064	
43	<i>ABO</i>	ABO blood group	rs8176746	9	136131322	
			rs8176719	9	136132909	
			rs11016116	10	129975450	
44	<i>MKI67</i>	Marker of proliferation Ki-67	rs148494166	10	129976030	
			rs115947774	10	130072795	
45	<i>RHOG</i>	Ras homolog family member G	rs138826089	11	3847190	monomorphic
46	<i>RRM1</i>	Ribonucleotide reductase M1	rs116472045	11	4111415	
47	<i>HBB</i>	Haemoglobin beta chain	rs334	11	5248232	
48	<i>TRIM5</i>	Tripartite motif containing 5	rs7935564	11	5718517	
49	<i>RTN3</i>	Reticulon 3	rs542998	11	63487386	
50	<i>GRIP1</i>	Glutamate receptor interacting protein 1	rs192909543	12	67366471	monomorphic
			rs1394263	12	67366537	
			rs1566830	12	67369898	
51	<i>CAND1</i>	Cullin-associated and neddylation-dissociated 1	rs12307123	12	67394950	
			rs10459266	12	67455888	
			rs2227507	12	68642647	
			rs1012356	12	68644618	
52	<i>IL22</i>	Interleukin 22	rs2227491	12	68646521	
			rs2227485	12	68647713	
			rs2227478	12	68648622	
53	<i>TPTE2</i>	Transmembrane phosphoinositide 3-phosphatase and tensin homolog 2	rs182873742	13	20050239	
54	<i>SPTB</i>	Spectrin, beta, erythrocytic	rs229587	14	65263300	
55	<i>LTBP2</i>	Latent transforming growth factor beta binding protein 2	rs74063230	14	75066093	monomorphic
56	<i>YLPM1</i>	YLP motif containing 1	rs10139016	14	75274288	
57	<i>RPS6KLI</i>	Ribosomal protein S6 kinase-like 1	rs3742785	14	75373034	
58	<i>ADCY9</i>	Adenylate cyclase 9	rs2230739	16	4033436	
			rs10775349	16	4079823	
59	<i>HBA</i>	Haemoglobin, alpha	α^+ thalassaemia	16	222846	
60	<i>IL4R</i>	Interleukin 4 receptor	rs1805015	16	27374180	
61	<i>ADORA2B</i>	Adenosine A2b receptor	rs2535611	17	15861332	
62	<i>NOS2</i>	Nitric oxide synthase 2A (inducible, hepatocytes)	rs2297518	17	26096597	
			rs1800482	17	26128509	

			rs9282799	17	26128728	
			rs8078340	17	26129212	
63	<i>BCAS3</i>	Microtubule associated cell migration factor	rs184142841	17	58855323	monomorphic
64	<i>TBX2</i>	T-box 2	rs73991577	17	59323072	
65	<i>EMRI</i>	ADGRE1 (Adhesion G-protein-coupled receptor E1)	rs373533	19	6919624	
			rs461645	19	6919753	
66	<i>ICAM</i>	Intercellular adhesion molecule 1	rs5498	19	10395683	
67	<i>GNAS</i>	Guanine nucleotide binding protein, alpha	rs8386	20	57485812	
68	<i>DERL3</i>	Derlin 3	rs1128127	22	24179132	
60	<i>CD40LG</i>	CD40 ligand (TNF superfamily, member 5)	rs3092945	X	135729609	
			rs1126535	X	135730555	
70	<i>G6PD</i>	Glucose-6-phosphate dehydrogenase	rs1050829	X	153763492	
			rs1050828	X	153764217	

Chr = chromosome, Removed in QC = SNPs that failed QC and were removed from further analysis.
Co-ordinate system is with respect to GRCh37.

Table S1: Summary of genes and genetic loci included in the study.

W3	iPLEX	rs148111931	ACGTTGGATGTGGTTCTTGGTAGGTGGG	ACGTTGGATGGAGCTGTGATGGAGTCACTG	106	98.4	68.8	53	52.4	dg	F	6864.5	gGGGTGATAGAGTGAAACCCCTG	A	7135.7	gGGGTGATAGAGTGAAACCCCTG	T	7191.6	gGGGTGATAGAGTGAAACCCCTG
W3	iPLEX	rs182873742	ACGTTGGATGGCTCTTGCTCATGTGATCTG	ACGTTGGATGAGGAATGAGCATAGGGAGAG	104	98.5	68.8	52	43.5	D	F	6927.5	TGTCATCTCCCTCAAACTAGACACC	C	7174.7	TGTCATCTCCCTCAAACTAGACACC	G	7214.7	TGTCATCTCCCTCAAACTAGACACC
W3	iPLEX	rs184895969	ACGTTGGATGAGGTCCTCCAAATTTTATC	ACGTTGGATGTGACCAATACACCTCCCAAC	100	90.2	68.8	47	30.4	D	R	7012.6	CAGTTTTTAATGACATCACATTG	T	7283.8	CAGTTTTTAATGACATCACATTG	C	7299.8	CAGTTTTTAATGACATCACATTG
W3	iPLEX	rs114726617	ACGTTGGATGTCCAATCAAGGAGCAGAG	ACGTTGGATGCTCTGTAAGCTCCAAGGCAT	119	94.6	68.8	50	30.4	DH	F	7079.6	AAAAAGTATTTGGGTACAACCA	A	7350.9	AAAAAGTATTTGGGTACAACCA	G	7366.9	AAAAAGTATTTGGGTACAACCA
W3	iPLEX	rs79691057	ACGTTGGATGGGCATACTACTCTCTCAC	ACGTTGGATGGGCATAGTGATGACCTGGAC	136	93.6	68.8	53	37.5	DA	R	7246.7	AACCTAGCATGACTTCATCTCAC	G	7493.9	AACCTAGCATGACTTCATCTCAC	A	7573.8	AACCTAGCATGACTTCATCTCAC
W3	iPLEX	rs74063230	ACGTTGGATGATCTTCCAGACTGGCTCC	ACGTTGGATGCCAGGCTGAGATAAATGAG	111	96.7	68.8	49	37.5	D	F	7473.9	CTGAGATAAAATGAGAAGTCTAGG	A	7745.1	CTGAGATAAAATGAGAAGTCTAGG	G	7761.1	CTGAGATAAAATGAGAAGTCTAGG
W3	iPLEX	rs10188961	ACGTTGGATGGCAATGTTTGAATTCAGAG	ACGTTGGATGATCACCTTACTCTCTCAC	95	86.4	68.8	48	24		R	7601	ACTTCAAAAATAACTGCAACTATTAC	G	7848.2	ACTTCAAAAATAACTGCAACTATTAC	A	7928.1	ACTTCAAAAATAACTGCAACTATTAC
W3	iPLEX	rs6674631	ACGTTGGATGATGTCTGTACTACTCTCC	ACGTTGGATGGGCTTTTGAATAAAGGGCAG	93	98.4	68.8	59	52	D	F	7624	TGGATTTGTAGAAGTGCCCTCCCA	A	7895.2	TGGATTTGTAGAAGTGCCCTCCCA	G	7911.2	TGGATTTGTAGAAGTGCCCTCCCA
W3	iPLEX	rs115947774	ACGTTGGATGATCCAATCAAGGATCATAG	ACGTTGGATGTCCAATGCAAGGAATGCC	119	86.7	68.8	56	42.3	D	R	7803.1	TAAGTAGATTAACCCCAATCTCAC	G	8050.3	TAAGTAGATTAACCCCAATCTCAC	A	8130.2	TAAGTAGATTAACCCCAATCTCAC
W3	iPLEX	rs2131263	ACGTTGGATGGCTCTGTATCTGAAACCAAC	ACGTTGGATGGCCTTACATGTAAGTACTCAC	128	90.3	68.8	50	28	DA	F	7946.2	gTGATCCAAAATTTAGAAAATCTCAC	C	8193.4	gTGATCCAAAATTTAGAAAATCTCAC	T	8273.3	gTGATCCAAAATTTAGAAAATCTCAC
W3	iPLEX	rs4316902	ACGTTGGATGGTCCCATGTTTATATAA	ACGTTGGATGGCCTTTTGAATAAAGGGCAG	118	80	68.8	45	15.4	D	F	8082.3	AGAAATTTAAAAGTAATGTGAAAAATA	A	8353.5	AGAAATTTAAAAGTAATGTGAAAAATA	G	8369.5	AGAAATTTAAAAGTAATGTGAAAAATA
W3	iPLEX	rs5598968	ACGTTGGATGGCAACAACCTCAGGAGTTG	ACGTTGGATGCTTTGGCCTCTCTGGATTG	133	95.7	68.8	46	18.5	DA	F	8237.4	GTTTTCTACCTTTAAAATGAATTAAT	A	8508.6	GTTTTCTACCTTTAAAATGAATTAAT	G	8524.6	GTTTTCTACCTTTAAAATGAATTAAT
W3	iPLEX	rs114726617	ACGTTGGATGGCTCTGTATCTGAAACCAAC	ACGTTGGATGGCCTTACATGTAAGTACTCAC	98	97.1	68.8	55	37	D	F	8300.4	GCTTCTGTCAGCAAGAAATAGAAAA	C	8547.6	GCTTCTGTCAGCAAGAAATAGAAAA	T	8627.5	GCTTCTGTCAGCAAGAAATAGAAAA
W3	iPLEX	rs186873296	ACGTTGGATGGTCCCATGTTTATATAA	ACGTTGGATGGCCTTTTGAATAAAGGGCAG	100	100	68.8	52	27.6	d	R	8862.8	AGAAATTTAAAAGTAATGTGAAAAATA	G	9110	AGAAATTTAAAAGTAATGTGAAAAATA	A	9189.9	AGAAATTTAAAAGTAATGTGAAAAATA
W3	iPLEX	rs144778284	ACGTTGGATGTGCACTTATAGCCCAATTC	ACGTTGGATGGGTAATAACAAAGGGTGG	107	97	68.8	60	44.8		R	8885.8	GGGTCCCTTTGCAAGGAACATCACAATT	T	9157	GGGTCCCTTTGCAAGGAACATCACAATT	C	9173	GGGTCCCTTTGCAAGGAACATCACAATT
W4	iPLEX	rs11016116	ACGTTGGATGTGAGAGGCACTCTGTACC	ACGTTGGATGGCTTTTATCTCACATGTGCC	119	98.6	18.5	46	60	D	R	4522	CACGTACCCGAGCAC	G	4769.1	CACGTACCCGAGCAC	A	4849.1	CACGTACCCGAGCAC
W4	iPLEX	rs8176750	ACGTTGGATGAACTGAGGCTTACTCTGG	ACGTTGGATGCAACCTCCCAAGAAAAATG	102	94.6	85	61	80	H	F	4578.9	GGCAGCCGCTCACGG	I	4866.1	GGCAGCCGCTCACGG	D	4906	GGCAGCCGCTCACGG
W4	iPLEX	rs144224092	ACGTTGGATGAACTGACCTCTGTGATCC	ACGTTGGATGGTGGTATCCTTTTACTG	136	82.3	18.5	47	60	DA	F	4633	CAGCACTTTGGGAGG	C	4880.2	CAGCACTTTGGGAGG	T	4960.1	CAGCACTTTGGGAGG
W4	iPLEX	rs10459266	ACGTTGGATGTCCAAAGTCCCTCCCTTTC	ACGTTGGATGTCCAAACAGAGGAGTGTAG	112	99.4	78.4	46	53.3	dh	R	4832.2	tCTCATCTTGGCAGCA	C	5119.4	tCTCATCTTGGCAGCA	A	5159.3	tCTCATCTTGGCAGCA
W4	iPLEX	rs192151845	ACGTTGGATGCACTCACTACTCTCTGCG	ACGTTGGATGTTAATGACAGACTGGAGGCG	112	96.6	18.5	55	68.8	D	R	5027.3	TTGGAGCAGGAGGCG	G	5274.5	TTGGAGCAGGAGGCG	A	5354.4	TTGGAGCAGGAGGCG
W4	iPLEX	rs7389579	ACGTTGGATGTGTCATCCCTGTGACTAAG	ACGTTGGATGGCTGGCTGTGAAAGATGAA	122	95.6	18.5	47	47.1	DA	F	5043.3	CTCCAACCTCACACAT	G	5330.5	CTCCAACCTCACACAT	T	5370.4	CTCCAACCTCACACAT
W4	iPLEX	rs56390333	ACGTTGGATGTCCAGAGCCCTGGCAG	ACGTTGGATGGAAGCTGAGGTTCACTGG	97	59.7	18.5	55	64.7	dh	R	5189.4	AGAACCACAGGCGGTC	G	5436.6	AGAACCACAGGCGGTC	C	5476.6	AGAACCACAGGCGGTC
W4	iPLEX	rs973128	ACGTTGGATGATATTGGAGGCTTAGCCAG	ACGTTGGATGGCCTTTTGTCTGCTGAG	107	98.4	18.5	55	55.6	Ds	R	5392.5	TTATCCCTTGCCCTGGCT	G	5639.7	TTATCCCTTGCCCTGGCT	C	5679.7	TTATCCCTTGCCCTGGCT
W4	iPLEX	rs148494166	ACGTTGGATGGGATCTGTTGAGCAGAAG	ACGTTGGATGATCTGGGAGATCTGCCCTG	111	98.1	18.5	46	47.1	d	F	5545.6	cCGTCTAGGTTTGGAA	C	5792.8	cCGTCTAGGTTTGGAA	T	5872.7	cCGTCTAGGTTTGGAA
W4	iPLEX	rs55868763	ACGTTGGATGTAGCCGTCGAAGTCTAGAT	ACGTTGGATGTCCACTCAGTCCCCCATC	101	96.1	18.5	52	52.6	DS	F	5723.7	CTCGCTGCCAGACTTCATAC	C	5970.9	CTCGCTGCCAGACTTCATAC	G	6010.9	CTCGCTGCCAGACTTCATAC
W4	iPLEX	rs4951074	ACGTTGGATGCAAGTCTATCTCTGCGG	ACGTTGGATGTCCAAACAGAGGAGTGTAG	100	100	18.5	46	42.1	D	R	5820.8	CTCTAGAGATAAAGCAGT	G	6068	CTCTAGAGATAAAGCAGT	A	6147.9	CTCTAGAGATAAAGCAGT
W4	iPLEX	rs186790584	ACGTTGGATGATCTCTGGAGAGCTTATG	ACGTTGGATGTGCCCTTTAGTGTCTCTAC	143	86.4	18.5	45	38.9	DA	F	5851.8	cTAAAGCAGCTGATTGAG	A	6123	cTAAAGCAGCTGATTGAG	T	6178.9	cTAAAGCAGCTGATTGAG
W4	iPLEX	rs76033371	ACGTTGGATGCCCAGTGTGTTAGCCTTTC	ACGTTGGATGCTCAGGCTTCTTAGTCTCC	132	96	18.5	46	40	DA	R	6026.9	TTCTAGTCTCCCTTAAACAAG	G	6274.1	TTCTAGTCTCCCTTAAACAAG	A	6354	TTCTAGTCTCCCTTAAACAAG
W4	iPLEX	rs3753036	ACGTTGGATGGCACTCACTACTCTCTGCG	ACGTTGGATGTCCAAACAGAGGAGTGTAG	120	98.4	78.4	47	44.4	d	R	6045	aaACCCACTCTAAGTGCTA	G	6292.1	aaACCCACTCTAAGTGCTA	A	6372.1	aaACCCACTCTAAGTGCTA
W4	iPLEX	rs12307123	ACGTTGGATGTCTGGAGAGACAATCAGAAC	ACGTTGGATGGCCAGTCAACTATCCCTG	114	97.8	18.5	46	40	d	F	6089	TGTTTATACTAGCTCTGGTC	C	6336.2	TGTTTATACTAGCTCTGGTC	T	6416.1	TGTTTATACTAGCTCTGGTC
W4	iPLEX	rs4650365	ACGTTGGATGTCAACATAAACAATATGAGC	ACGTTGGATGCTGACAGTCTATAGAGTCCC	108	87.5	18.5	45	50	Ds	F	6317.1	tgGCTGAACCTCCTACTACT	C	6564.3	tgGCTGAACCTCCTACTACT	T	6644.2	tgGCTGAACCTCCTACTACT
W4	iPLEX	rs14155199	ACGTTGGATGCAAGTCTACTGCACTTATAA	ACGTTGGATGTAGAGGGAATTTGTCTAAG	108	82.8	18.5	49	42.9	dh	F	6445.2	GTCTAGCTACATAGAAGCTGT	C	6692.4	GTCTAGCTACATAGAAGCTGT	G	6732.4	GTCTAGCTACATAGAAGCTGT
W4	iPLEX	rs79100774	ACGTTGGATGCCCCCTGGATGGTTATATTC	ACGTTGGATGTAGAAGAATGATTGGTGG	121	61.8	18.5	48	38.1	dA	F	6509.3	ATTGGTGGCTATAGCATAAAG	C	6756.4	ATTGGTGGCTATAGCATAAAG	G	6796.5	ATTGGTGGCTATAGCATAAAG
W4	iPLEX	rs184908374	ACGTTGGATGGTGGCAATTTGTTGGTTAC	ACGTTGGATGCACAACCTCTGGTGTGAATG	100	97.1	18.5	51	38.1		R	6773.4	tTGGTGTGAATGCACCAATAT	G	7020.6	tTGGTGTGAATGCACCAATAT	A	7100.5	tTGGTGTGAATGCACCAATAT
W4	iPLEX	rs149914432	ACGTTGGATGAAAGTACTGCCACTGCCAAC	ACGTTGGATGAGCAGAGCAAGTTGGTAG	131	94.7	18.5	49	31.8	DAs	F	6873.5	AAGTTGGTAGCAAAAAAGAGAAC	C	7120.7	AAGTTGGTAGCAAAAAAGAGAAC	A	7144.7	AAGTTGGTAGCAAAAAAGAGAAC
W4	iPLEX	rs151293197	ACGTTGGATGGCAGAGTCCAGATTACAC	ACGTTGGATGGATTTTCTGTCTCAAGC	101	87.9	18.5	52	38.1	d	F	7047.6	ggTGTCTCAAGCAAATGCACAT	C	7294.8	ggTGTCTCAAGCAAATGCACAT	G	7334.8	ggTGTCTCAAGCAAATGCACAT
W4	iPLEX	rs72933310	ACGTTGGATGCCTCTATGCTTTTCTATGGC	ACGTTGGATGGTAGGATATCTATGCAATGG	96	92.9	18.5	46	30.4	D	F	7070.6	GATATCTATGCAATGGAATACTA	C	7317.8	GATATCTATGCAATGGAATACTA	T	7397.7	GATATCTATGCAATGGAATACTA
W4	iPLEX	rs6457452	ACGTTGGATGAAACGGCCAGCCTGAGGAG	ACGTTGGATGGTTCGCTCTGGAAAGCCTG	107	92.8	18.5	54	50	s	R	7081.6	IGGAGTCACTCTGAAAGACGAAA	T	7352.8	IGGAGTCACTCTGAAAGACGAAA	C	7368.8	IGGAGTCACTCTGAAAGACGAAA
W4	iPLEX	rs72933304	ACGTTGGATGGGAAAGCTATCCTCAAGGTG	ACGTTGGATGATCTGTTCTGCCCTCAC	137	94.7	18.5	58	50	DA	F	7174.7	GTTTCTGCCCTTACCCTTACAA	C	7421.9	GTTTCTGCCCTTACCCTTACAA	A	7445.9	GTTTCTGCCCTTACCCTTACAA
W4	iPLEX	rs28459062	ACGTTGGATGAGTCTAGTTCATGGAGAGGG	ACGTTGGATGATCCAGTCCCCTCACTTTC	108	98.3	18.5	47	30.4	D	R	7236.7	aCCATCACTTCAATGTTATCTATA	C	7507.9	aCCATCACTTCAATGTTATCTATA	C	7523.9	aCCATCACTTCAATGTTATCTATA
W4	iPLEX	rs1394263	ACGTTGGATGCTCATCAGAGTAAATGCTC	ACGTTGGATGTGGACCTTAGCTGAGTGTG	123	96.5	18.5	46	26.1	DA	R	7463.9	gTGAATTAGAATGCATTAGTAAGAA	T	7735.1	gTGAATTAGAATGCATTAGTAAGAA	C	7751.1	gTGAATTAGAATGCATTAGTAAGAA
W4	iPLEX	rs13103597	ACGTTGGATGCCCTGCAAACTCTGTCAATC	ACGTTGGATGAAGGGCTACAAAGTTATGG	117	97.4	18.5	46	24		R	7654	TACAAACGTTATGGTTTTAAATATC	T	7925.2	TACAAACGTTATGGTTTTAAATATC	C	7941.2	TACAAACGTTATGGTTTTAAATATC
W4	iPLEX	rs184142841	ACGTTGGATGGTTTCTTTAGTACCCTCC	ACGTTGGATGTCCAAAGAGACATACGG	108	92.8	61.8	46	20	D	R	7656	ATGAACACATAAAAAATGTTTCA	G	7903.2	ATGAACACATAAAAAATGTTTCA	A	7983.1	ATGAACACATAAAAAATGTTTCA
W4	iPLEX	rs75180423	ACGTTGGATGTGCCATCCAGATGGCAAAC	ACGTTGGATGGGAAACCTTACTCTAGTGA	116	93.5	18.5	45	24	d	R	7672	GATAATTAGTGTCTTACAAATAAA	C	7959.2	GATAATTAGTGTCTTACAAATAAA	A	7999.1	GATAATTAGTGTCTTACAAATAAA
W4	iPLEX	rs1541255	ACGTTGGATGTAGCCGTCCGAAGTCTAGAT	ACGTTGGATGTCCACTCAGTCCCCCATC	101	96.1	18.5	57	50	DSH	R	7848.1	CCGTCCGAAGTCTAGATCTTCTTCC	G	8095.3	CCGTCCGAAGTCTAGATCTTCTTCC	T	8119.3	CCGTCCGAAGTCTAGATCTTCTTCC
W4	iPLEX	rs142712208	ACGTTGGATGTTTCTTCTCTGCAGACAC	ACGTTGGATGGTGTACTAACAATACCAG	129	86.2	18.5	45	18.5	A	F	8295.4	CCATTATGATAATTGATGATTAATAAT	A	8566.7	CCATTATGATAATTGATGATTAATAAT	G	8582.7	CCATTATGATAATTGATGATTAATAAT
W4	iPLEX	rs114980857	ACGTTGGATGGTCCACTTTCTAGGTCC	ACGTTGGATGGAGTCCAGGCTGATAAGGTC	85	96.3	18.5	56	44.4	d	F	8421.5	GATAAGGTTTCACATGGAGATGAGGAA	C	8668.7	GATAAGGTTTCACATGGAGATGAGGAA	T	8748.6	GATAAGGTTTCACATGGAGATGAGGAA
W4	iPLEX	rs75731597	ACGTTGGATGGTTCACATAAGATATGACTG	ACGTTGGATGTGCTTCCATCACAACCTG	113	85.3	85.3	45	14.3		F	8547.6	TGTTTTAATTAATTTCTCTGAAATTA	C	8794.8	TGTTTTAATTAATTTCTCTGAAATTA	A	8818.8	TGTTTTAATTAATTTCTCTGAAATTA
W4	iPLEX	rs74954675	ACGTTGGATGTTAAGGCATGCAGAGGTTG	ACGTTGGATGTACATTTTAAAGAGAG	135	66.3	18.5	47	21.4	dA	R	8611.7	GAAAAATCTTTAAATGATCACAATG	C	8898.9	GAAAAATCTTTAAATGATCACAATG	A	8938.8	GAAAAATCTTTAAATGATCACAATG

Assay definitions for the Agena iPLEX Mass-Array genotyping platform for the 129 SNPs investigated in this study. The designs were undertaken using the MassARRAY® Assay Design 3.1 software, which grouped the assays into 4 multiplexes assigned W1 - W4 (WELL column). Field definitions are as per the Agena assay design software users' guide (MassARRAY® Assay Design 3.1, Sequenom Inc., San Diego, Ca, USA).

Table S2: Assay definitions for the Agena iPLEX genotyping platform.

Gene	SNPID	Alleles (ancestral / derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	MO	Unadjusted [§]				Adjusted [§]			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.08	3940	All	SM	0.07	2215	A	0.95	0.82	1.09	0.4382	0.92	0.79	1.06	0.4900
AJAP1	rs6674631	G/A	1	0.13	3928	All	SM	0.12	2212	A	0.89	0.79	1.00	0.0434	0.88	0.78	1.00	0.04
ATP2B4	rs10900585	G/T	1	0.34	3762	All	SM	0.33	2200	R	1.29	1.07	1.56	0.0056	1.23	1.10	1.81	0.0090
ATP2B4	rs1541255	A/G	1	0.33	3933	All	SM	0.31	2211	R	0.74	0.62	0.89	0.0012	0.76	0.63	0.92	0.001
ATP2B4	rs3753036	G/A	1	0.04	3939	All	SM	0.04	2217	R	1.60	0.42	6.07	0.4940	1.25	0.28	5.49	0.8400
ATP2B4	rs4951074	G/A	1	0.32	3934	All	SM	0.31	2215	R	0.77	0.64	0.93	0.0068	0.77	0.65	0.94	0.007
ATP2B4	rs55868763	G/C	1	0.33	3931	All	SM	0.31	2209	R	0.74	0.62	0.89	0.0013	0.75	0.61	0.92	0.0010
CR1	rs17047660	A/G	1	0.16	3923	All	SM	0.17	2228	R	1.24	0.90	1.72	0.1932	1.14	0.81	1.61	0.18
CR1	rs17047661	G/A	1	0.32	3934	All	SM	0.33	2223	A	1.07	0.98	1.15	0.1249	1.06	0.98	1.16	0.1300
DARC	rs2814778	G/A	1	0.00	3944	All	SM	0.00	1933	H	0.83	0.33	2.10	0.6934	0.77	0.30	1.97	0.69
GBP7	rs1803632	G/C	1	0.49	3942	All	SM	0.51	2230	R	1.11	0.98	1.25	0.0940	1.10	0.96	1.25	0.1200
IL10	rs1800890	A/T	1	0.24	3945	All	SM	0.23	2233	R	0.69	0.55	0.88	0.0025	0.72	0.56	0.93	0.002
IL10	rs1800896	T/C	1	0.38	3943	All	SM	0.36	2225	R	0.90	0.77	1.05	0.1961	0.91	0.77	1.07	0.2200
IL10	rs3024500	G/A	1	0.45	3928	All	SM	0.47	2226	R	1.18	1.04	1.34	0.0129	1.17	1.02	1.33	0.01
LPHN2	rs146428334	A/G	1	0.00	3930	All	SM	0.00	2216	H	1.29	0.74	2.23	0.3751	1.30	0.74	2.30	0.3600
LPHN2	rs4650365	T/C	1	0.20	3936	All	SM	0.19	2215	H	0.85	0.76	0.95	0.0050	0.86	0.76	0.97	0.003
LPHN2	rs72933304	C/A	1	0.09	3942	All	SM	0.07	2216	A	0.80	0.69	0.92	0.0019	0.83	0.72	0.96	0.0020
LPHN2	rs72933310	T/C	1	0.09	3927	All	SM	0.08	2214	A	0.85	0.74	0.97	0.0158	0.87	0.75	0.98	0.01
LPHN2	rs72933350	T/C	1	0.05	3940	All	SM	0.04	2215	D	0.75	0.62	0.92	0.0037	0.82	0.67	0.94	0.0040
IL1A	rs17561	G/T	2	0.16	3938	All	SM	0.18	2232	R	1.47	1.08	2.00	0.0152	1.49	1.08	2.05	0.01
IL1B	rs1143634	C/T	2	0.11	3941	All	SM	0.13	2229	A	1.08	0.96	1.21	0.1850	1.06	0.94	1.19	0.2300
LAPTM4A	rs973128	C/G	2	0.44	3885	All	SM	0.44	2183	H	1.03	0.92	1.14	0.6460	0.98	0.87	1.11	0.68
LOC727982	rs10188961	A/G	2	0.40	3934	All	SM	0.42	2211	D	1.21	1.08	1.35	0.0009	1.14	1.05	1.24	0.0007
LOC727982	rs1371474	T/C	2	0.27	3933	All	SM	0.29	2216	H	1.29	1.16	1.43	4.19×10 ⁻⁶	1.34	1.19	1.50	3.86×10 ⁻⁶
LOC727982	rs1371478	C/T	2	0.27	3909	All	SM	0.29	2209	H	1.30	1.17	1.45	1.90×10 ⁻⁶	1.35	1.20	1.51	2.31×10 ⁻⁶
SDC1	rs11899121	C/G	2	0.47	3921	All	SM	0.47	2210	H	1.06	0.96	1.18	0.2570	1.06	0.95	1.19	0.23
ZSWIM2	rs144778284	C/T	2	0.00	3937	All	SM	0.00	2216	R	-	-	-	-	-	-	-	-
ZSWIM2	rs4316902	A/G	2	0.18	3910	All	SM	0.17	2205	R	0.79	0.57	1.08	0.1344	0.81	0.58	1.14	0.12
ARL14	rs74954675	A/C	3	0.08	3907	All	SM	0.10	2200	H	1.24	1.07	1.43	0.0039	1.22	1.05	1.42	0.0030
ARL14	rs75731597	A/C	3	0.08	3491	All	SM	0.10	2173	H	1.27	1.10	1.47	0.0013	1.25	1.07	1.46	0.001
ARL14	rs76033371	A/G	3	0.09	3939	All	SM	0.10	2207	H	1.22	1.06	1.40	0.0060	1.19	1.03	1.38	0.0060
B3GALNT1	rs12107243	G/C	3	0.49	3934	All	SM	0.49	2213	D	0.95	0.84	1.08	0.4393	0.97	0.85	1.10	0.45
IL17RD	rs6780995	A/G	3	0.41	3939	All	SM	0.40	2230	R	0.90	0.78	1.03	0.1283	0.91	0.79	1.06	0.1700
IL17RE	rs708567	A/G	3	0.50	3881	All	SM	0.50	2221	R	0.97	0.86	1.10	0.6779	0.98	0.86	1.12	0.66
TLR9	rs187084	T/C	3	0.33	3933	All	SM	0.32	1937	A	0.95	0.88	1.04	0.2620	0.92	0.84	1.00	0.2200
FREM3	rs149914432	A/C	4	0.10	3915	All	SM	0.06	2202	A	0.60	0.51	0.69	8.57×10 ⁻¹³	0.64	0.53	0.78	6.20×10 ⁻¹³
FREM3	rs184895969	C/T	4	0.00	3940	All	SM	0.00	2217	H	0.83	0.45	1.53	0.5508	1.00	0.51	1.94	0.5600
FREM3	rs184908374	G/A	4	0.00	3943	All	SM	0.00	2217	H	0.95	0.50	1.82	0.8842	1.14	0.56	2.30	0.9
FREM3	rs186790584	A/T	4	0.10	3941	All	SM	0.06	2217	A	0.59	0.50	0.68	2.71×10 ⁻¹³	0.60	0.51	0.7	2.24×10 ⁻¹³
FREM3	rs186873296	A/G	4	0.10	3926	All	SM	0.06	2206	A	0.57	0.49	0.66	4.23×10 ⁻¹⁴	0.64	0.53	0.79	3.18×10 ⁻¹⁴
GAB1	rs7663712	A/G	4	0.02	3938	All	SM	0.03	2216	H	1.09	0.86	1.39	0.4862	1.04	0.81	1.31	0.4830
INPP4B	rs13103597	C/T	4	0.27	3925	All	SM	0.23	2215	A	0.82	0.75	0.89	7.41×10 ⁻⁶	0.89	0.80	0.98	6.29×10 ⁻⁶
INPP4B	rs77389579	G/T	4	0.07	3937	All	SM	0.04	2209	A	0.61	0.51	0.72	4.48×10 ⁻⁹	0.64	0.53	0.77	4.79×10 ⁻⁹
TLR1	rs4833095	C/T	4	0.09	3921	All	SM	0.10	1936	R	1.49	0.87	2.57	0.1519	1.26	0.70	2.26	0.15
TLR6	rs5743809	T/C	4	0.06	3884	All	SM	0.06	1847	R	1.41	0.62	3.21	0.4159	1.52	0.65	3.56	0.4120
TLR6	rs5743810	C/T	4	0.02	3936	All	SM	0.01	1940	R	-	-	-	-	-	-	-	-
USP38	rs28459062	T/C	4	0.16	3931	All	SM	0.18	2217	A	1.14	1.03	1.26	0.0095	1.10	1.10	1.27	0.0100
USP38	rs4266246	C/T	4	0.22	3928	All	SM	0.22	2212	R	1.09	0.86	1.40	0.4724	1.02	0.79	1.31	0.416
C6	rs1801033	A/C	5	0.47	3927	All	SM	0.47	1933	A	0.99	0.92	1.07	0.8171	0.99	0.91	1.07	0.8230
IL13	rs20541	C/T	5	0.22	3796	All	SM	0.21	1850	R	0.88	0.67	1.14	0.3288	0.86	0.65	1.14	0.39
IRF1	rs2706384	C/A	5	0.38	3819	All	SM	0.36	1886	A	0.95	0.87	1.03	0.1835	0.94	0.86	1.02	0.2140
CTL4	rs2242665	A/G	6	0.26	3853	All	SM	0.27	1911	R	1.16	0.94	1.43	0.1737	1.15	0.92	1.43	0.17
GABBR1	rs192151845	G/A	6	0.01	3943	All	SM	0.01	2212	H	1.07	0.69	1.66	0.7515	1.07	0.68	1.69	0.8300
HCG4	rs114980857	C/T	6	0.21	3843	All	SM	0.22	2188	H	1.04	0.93	1.17	0.4713	1.01	0.89	1.14	0.495
HSPA1B	rs6457452	C/T	6	0.19	3917	All	SM	0.18	2212	D	0.88	0.79	0.98	0.0243	0.88	0.78	0.99	0.0200
IL20RA	rs1555498	C/T	6	0.46	3947	All	SM	0.49	2228	A	1.05	0.98	1.13	0.1810	1.06	0.98	1.14	0.179
LTA	rs2239704	G/T	6	0.12	3896	All	SM	0.13	2187	D	1.07	0.95	1.21	0.2709	1.03	0.90	1.18	0.3110
LTA	rs909253	T/C	6	0.49	3921	All	SM	0.49	2216	H	1.03	0.92	1.14	0.6090	1.06	0.95	1.19	0.501
PLEKHG1	rs114726617	G/A	6	0.04	3940	All	SM	0.04	2217	A	1.15	0.95	1.39	0.1474	1.20	0.98	1.47	0.1720
PLEKHG1	rs141555199	C/G	6	0.02	3943	All	SM	0.02	2218	R	0.00	0.00	-	0.1424	-	-	-	-
PLEKHG1	rs142712208	G/A	6	0.03	3927	All	SM	0.04	2218	A	1.18	0.96	1.44	0.1147	1.27	1.02	1.57	0.1160
PLEKHG1	rs15116938	C/T	6	0.03	3939	All	SM	0.04	2216	A	1.13	0.93	1.39	0.2207	1.23	0.99	1.52	0.225
PLEKHG1	rs151293197	C/G	6	0.03	3940	All	SM	0.04	2217	A	1.12	0.91	1.37	0.2790	1.19	0.96	1.48	0.2850
PLEKHG1	rs2131263	C/T	6	0.04	3938	All	SM	0.04	2216	A	1.14	0.95	1.38	0.1629	1.20	0.98	1.46	0.172
PLEKHG1	rs55958968	A/G	6	0.05	3932	All	SM	0.06	2217	H	1.20	1.01	1.43	0.0376	1.21	1.01	1.45	0.0300
PLEKHG1	rs76924464	G/A	6	0.03	3940	All	SM	0.04	2217	A	1.12	0.92	1.37	0.2673	1.20	0.97	1.48	0.273
PLEKHG1	rs79100774	G/C	6	0.03	3939	All	SM	0.04	2217	A	1.11	0.91	1.36	0.2934	1.20	0.97	1.48	0.2980
SNORD48	rs116288147	C/T	6	0.24	3896	All	SM	0.23	2191	D	0.93	0.83	1.03	0.1621	0.92	0.82	1.04	0.138
TNF	rs1799964	T/C	6	0.27	3938	All	SM	0.27	2204	H	0.95	0.86	1.06	0.3902	0.94	0.83	1.05	0.4240
TNF	rs1800629	G/A	6	0.09	3940	All	SM	0.09	2230	H	0.92	0.80	1.06	0.2422	0.92	0.80	1.07	0.286
TNF	rs1800750	G/A	6	0.05	3868	All	SM	0.05	1281	A	0.94	0.77	1.16	0.5740	0.93	0.74	1.17	0.5100
TNF	rs3093662	A/G	6	0.11	3936	All	SM	0.11	2219	R	0.75	0.45	1.23	0.2424	0.74	0.45	1.23	0.247
TNF	rs361525	G/A	6	0.07	3948	All	SM	0.07	2224	R	0.66	0.29	1.4					

<i>MKI67</i>	rs148494166	T/C	10	0.00	3913	All	SM	0.00	2137	H	1.36	0.63	2.91	0.4401	1.16	0.53	2.56	0.427
<i>MKI67</i>	rs11016116	A/G	10	0.02	3944	All	SM	0.02	2218	R	1.65	0.41	6.68	0.4877	1.84	0.42	8.00	0.4810
<i>MKI67</i>	rs115947774	A/G	10	0.00	3931	All	SM	0.00	2206	H	0.49	0.23	1.06	0.0564	0.48	0.22	1.05	0.05
<i>HBB</i>	rs334	A/T	11	0.08	3949	All	SM	0.02	2227	H	0.14	0.12	0.21	6.81×10 ⁻⁵⁹	0.15	0.11	0.20	2.61×10 ⁻⁵⁸
<i>RRM1</i>	rs116472045	A/G	11	0.02	3939	All	SM	0.02	2213	R	-	-	-	-	-	-	-	-
<i>RTN3</i>	rs542998	T/C	11	0.43	3929	All	SM	0.44	1911	D	1.06	0.94	1.20	0.3118	1.09	0.96	1.23	0.2660
<i>TRIM5</i>	rs7935564	G/A	11	0.44	3871	All	SM	0.46	1930	A	1.08	1.00	1.17	0.0473	1.00	0.92	1.09	0.044
<i>CAND1</i>	rs10459266	C/A	12	0.21	3938	All	SM	0.22	2217	D	1.14	1.02	1.27	0.0179	1.13	1.01	1.27	0.0130
<i>CAND1</i>	rs12307123	C/T	12	0.23	3933	All	SM	0.24	2216	A	1.09	1.00	1.19	0.0566	1.11	0.99	1.24	0.04
<i>CAND1</i>	rs1566830	G/T	12	0.26	3940	All	SM	0.28	2215	A	1.11	1.02	1.20	0.0188	1.09	1.00	1.20	0.0150
<i>GRIP1</i>	rs1394263	T/C	12	0.26	3914	All	SM	0.28	2205	A	1.11	1.02	1.21	0.0150	1.10	1.01	1.20	0.012
<i>IL22</i>	rs1012356	A/T	12	0.49	3940	All	SM	0.48	2225	H	1.08	0.97	1.20	0.1519	1.07	0.96	1.20	0.1480
<i>IL22</i>	rs2227478	A/G	12	0.38	3939	All	SM	0.39	2230	R	0.97	0.84	1.13	0.7107	0.95	0.81	1.11	0.688
<i>IL22</i>	rs2227485	G/A	12	0.42	3928	All	SM	0.41	2221	R	0.86	0.75	0.99	0.0402	0.84	0.72	0.97	0.0370
<i>IL22</i>	rs2227491	C/T	12	0.40	3870	All	SM	0.42	2192	D	1.09	0.98	1.22	0.1196	1.07	0.95	1.20	0.1
<i>IL22</i>	rs2227507	T/C	12	0.04	3948	All	SM	0.04	2235	A	0.96	0.80	1.16	0.6840	0.96	0.79	1.17	0.7360
<i>TPTE2</i>	rs182873742	C/G	13	0.00	3940	All	SM	0.00	2202	H	4.92	0.50	48.34	0.1362	4.92	0.40	60.10	0.133
<i>RPS6KL1</i>	rs3742785	C/A	14	0.29	3714	All	SM	0.30	2115	H	1.20	1.08	1.34	0.0010	1.18	1.05	1.33	0.0016
<i>SPTB</i>	rs229587	T/C	14	0.34	3872	All	SM	0.34	2167	A	1.02	0.94	1.10	0.6677	1.01	0.93	1.09	0.637
<i>YLFM1</i>	rs10139016	C/T	14	0.25	3937	All	SM	0.27	2214	D	1.12	1.01	1.25	0.0313	1.11	0.99	1.24	0.0300
<i>ADCY9</i>	rs10775349	C/G	16	0.20	3941	All	SM	0.20	1937	R	1.09	0.82	1.44	0.5625	1.11	0.83	1.50	0.614
<i>ADCY9</i>	rs2230739	A/G	16	0.09	3944	All	SM	0.08	1940	H	0.87	0.74	1.01	0.0689	0.83	0.70	0.97	0.0500
<i>HBA</i>	α ^{-3.7} -thalassaemia	I/D	16	0.41	3946	All	SM	0.36	2143	A	0.82	0.76	0.89	6.59×10 ⁻⁷	0.83	0.76	0.90	2.06×10 ⁻⁶
<i>IL4</i>	rs2243250	T/C	16	0.20	3902	All	SM	0.22	2205	H	1.11	0.99	1.24	0.0632	1.09	0.97	1.23	0.0500
<i>IL4R</i>	rs1805015	T/C	16	0.42	3927	All	SM	0.43	2228	H	1.05	0.94	1.17	0.3772	1.05	0.94	1.18	0.368
<i>ADORA2B</i>	rs2535611	T/C	17	0.10	3911	All	SM	0.11	1935	A	1.08	0.96	1.23	0.2005	1.06	0.93	1.21	0.2020
<i>NOS2</i>	rs1800482	G/C	17	0.08	3922	All	SM	0.07	2220	A	0.97	0.84	1.12	0.6718	0.95	0.82	1.10	0.581
<i>NOS2</i>	rs2297518	G/A	17	0.14	3910	All	SM	0.13	1924	D	0.95	0.84	1.08	0.4227	0.96	0.84	1.10	0.4430
<i>NOS2</i>	rs8078340	C/T	17	0.21	3948	All	SM	0.21	2233	R	0.82	0.62	1.09	0.1624	0.78	0.58	1.04	0.146
<i>NOS2</i>	rs9282799	C/T	17	0.03	3949	All	SM	0.03	2224	H	1.14	0.91	1.44	0.2515	1.13	0.88	1.44	0.2990
<i>TBX2</i>	rs73991577	A/G	17	0.00	3937	All	SM	0.00	2215	H	0.48	0.19	1.20	0.0976	0.52	0.20	1.34	0.1
<i>EMR1</i>	rs373533	G/T	19	0.46	3863	All	SM	0.49	1927	A	1.09	1.01	1.18	0.0345	1.07	0.99	1.17	0.0400
<i>EMR1</i>	rs461645	C/T	19	0.47	3933	All	SM	0.49	1939	A	1.09	1.00	1.17	0.0374	1.07	0.99	1.16	0.048
<i>ICAM</i>	rs5498	A/G	19	0.11	3908	All	SM	0.12	2197	H	1.05	0.93	1.20	0.4327	1.03	0.89	1.18	0.4550
<i>GNAS</i>	rs8386	C/T	20	0.19	3944	All	SM	0.19	2229	H	0.91	0.81	1.02	0.1060	0.92	0.82	1.04	0.11
<i>DERL3</i>	rs1128127	G/A	22	0.50	3911	All	SM	0.47	2226	A	0.94	0.88	1.02	0.1241	0.93	0.86	1.01	0.1250
<i>CD40LG</i>	rs1126535	T/C	X	0.15	3934	M+F	SM	0.16	2230	H	0.86	0.72	1.02	0.0843	0.85	0.71	1.03	0.09
<i>CD40LG</i>	rs1126535	T/C	X	0.16	1947	F	SM	0.14	1080	A	0.85	0.73	0.99	0.0325	0.87	0.76	1.00	0.0425
<i>CD40LG</i>	rs1126535	C/T	X	0.85	1987	M	SM	0.83	1150	HM	0.84	0.69	1.03	0.0903	0.85	0.70	1.04	0.100257
<i>CD40LG</i>	rs3092945	T/C	X	0.21	3927	M+F	SM	0.26	2226	A	1.14	1.07	1.23	0.0002	1.15	1.06	1.23	0.0002
<i>CD40LG</i>	rs3092945	T/C	X	0.21	1942	F	SM	0.25	1082	A	1.25	1.10	1.42	0.0004	1.28	1.13	1.45	0.0002
<i>CD40LG</i>	rs3092945	C/T	X	0.78	1985	M	SM	0.74	1144	HM	0.83	0.70	0.99	0.0338	0.80	0.68	0.95	0.0100
<i>G6PD</i>	rs1050828	C/T	X	0.19	3940	M+F	SM	0.19	2220	H	0.82	0.69	0.96	0.0153	0.80	0.68	0.96	0.015
<i>G6PD</i>	rs1050828	C/T	X	0.20	1951	F	SM	0.18	1074	H	0.82	0.70	0.97	0.0100	0.81	0.69	0.96	0.0100
<i>G6PD</i>	rs1050828	C/T	X	0.19	1989	M	SM	0.20	1146	HM	1.11	0.92	1.34	0.2686	1.08	0.90	1.30	0.36
<i>G6PD</i>	rs1050829	T/C	X	0.40	3928	M+F	SM	0.42	2221	H	0.85	0.73	0.99	0.0342	0.86	0.73	1.01	0.0540
<i>G6PD</i>	rs1050829	T/C	X	0.41	1952	F	SM	0.41	1075	H	0.85	0.73	0.99	0.0358	0.86	0.74	1.00	0.04581
<i>G6PD</i>	rs1050829	C/T	X	0.60	1976	M	SM	0.58	1146	HM	0.90	0.77	1.04	0.1486	0.93	0.79	1.05	0.1526

* The rs176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and rs334 genotype.

Table S3: Odds ratios for all SM for all polymorphisms investigated.

Gene	SNPID	Alleles (ancestral/ derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	MO	Unadjusted				Adjusted ^s			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.08	3940	All	CM	0.07	1231	H	0.94	0.77	1.13	0.5004	0.92	0.76	1.12	0.5601
AJAP1	rs6674631	G/A	1	0.13	3928	All	CM	0.11	1225	A	0.83	0.72	0.96	0.0100	0.83	0.71	0.96	0.01161
ATP2B4	rs10900585	G/T	1	0.34	3762	All	CM	0.32	1223	R	1.21	1.06	1.54	0.0210	1.39	0.89	2.18	0.0187
ATP2B4	rs1541255	A/G	1	0.33	3933	All	CM	0.30	1226	R	0.68	0.54	0.86	0.0011	0.69	0.54	0.89	0.0011
ATP2B4	rs3753036	G/A	1	0.04	3939	All	CM	0.04	1229	R	2.16	0.50	9.27	0.3208	1.51	0.28	8.19	0.7274
ATP2B4	rs4951074	G/A	1	0.32	3934	All	CM	0.30	1229	R	0.73	0.58	0.93	0.0092	0.97	0.60	1.55	0.0084671
ATP2B4	rs55868763	G/C	1	0.33	3931	All	CM	0.30	1227	R	0.68	0.54	0.87	0.0011	0.67	0.52	0.86	0.0009
CR1	rs17047660	A/G	1	0.16	3923	All	CM	0.18	1221	A	1.14	1.01	1.29	0.0397	1.11	0.98	1.26	0.058625
CR1	rs17047661	G/A	1	0.32	3934	All	CM	0.34	1217	A	1.10	1.00	1.21	0.0614	1.12	1.01	1.24	0.0732
DARC	rs2814778	G/A	1	0.00	3944	All	CM	0.00	1093	H	0.80	0.26	2.49	0.6942	0.74	0.24	2.34	0.69406
GBP7	rs1803632	G/C	1	0.49	3942	All	CM	0.51	1223	A	1.07	0.98	1.17	0.1484	1.08	0.98	1.18	0.1564
IL10	rs1800890	A/T	1	0.24	3945	All	CM	0.23	1225	R	0.74	0.55	0.99	0.0387	0.76	0.55	0.99	0.040069
IL10	rs1800896	T/C	1	0.38	3943	All	CM	0.36	1221	R	0.87	0.72	1.06	0.1673	0.87	0.71	1.06	0.1683
IL10	rs3024500	G/A	1	0.45	3928	All	CM	0.49	1222	R	1.26	1.08	1.47	0.0041	1.24	1.06	1.46	0.0083114
LPHN2	rs146428334	A/G	1	0.00	3930	All	CM	0.01	1230	H	1.51	0.80	2.86	0.2188	1.51	0.79	2.90	0.2102
LPHN2	rs4650365	T/C	1	0.20	3936	All	CM	0.19	1227	H	0.83	0.72	0.96	0.0091	0.84	0.72	0.97	0.0064098
LPHN2	rs72933304	C/A	1	0.09	3942	All	CM	0.07	1230	A	0.82	0.69	0.98	0.0279	0.86	0.71	0.99	0.0371
LPHN2	rs72933310	T/C	1	0.09	3927	All	CM	0.08	1229	D	0.86	0.72	1.03	0.1014	0.90	0.75	1.08	0.10928
LPHN2	rs72933350	T/C	1	0.05	3940	All	CM	0.04	1229	D	0.76	0.60	0.96	0.0203	0.82	0.64	1.05	0.0245
IL1A	rs17561	G/T	2	0.16	3938	All	CM	0.18	1224	R	1.40	0.96	2.04	0.0860	1.43	0.97	2.11	0.084987
IL1B	rs1143634	C/T	2	0.11	3941	All	CM	0.12	1221	H	1.07	0.91	1.25	0.4032	1.06	0.90	1.25	0.4466
LAPTM4A	rs973128	C/G	2	0.44	3885	All	CM	0.45	1214	D	1.08	0.93	1.24	0.3029	1.02	0.88	1.18	0.28849
LOC727982	rs10188961	A/G	2	0.40	3934	All	CM	0.41	1225	D	1.14	1.00	1.31	0.0551	1.17	1.01	1.35	0.0526
LOC727982	rs1371474	T/C	2	0.27	3933	All	CM	0.29	1230	H	1.32	1.16	1.50	4.21×10 ⁻⁵	1.37	1.20	1.58	5.97×10 ⁻⁵
LOC727982	rs1371478	C/T	2	0.27	3909	All	CM	0.29	1226	H	1.31	1.15	1.50	4.61×10 ⁻⁵	1.37	1.19	1.57	7.19×10 ⁻⁵
SDC1	rs11899121	C/G	2	0.47	3921	All	CM	0.48	1227	H	1.08	0.95	1.23	0.2659	1.06	0.93	1.21	0.22618
ZSWIM2	rs144778284	C/T	2	0.00	3937	All	CM	0.00	1229	R	-	-	-	-	-	-	-	-
ZSWIM2	rs4316902	A/G	2	0.18	3910	All	CM	0.17	1225	R	0.71	0.47	1.08	0.0984	0.76	0.50	1.17	0.10588
ARL14	rs74954675	A/C	3	0.08	3907	All	CM	0.09	1219	H	1.23	1.03	1.46	0.0227	1.24	1.03	1.49	0.0167
ARL14	rs75731597	A/C	3	0.08	3491	All	CM	0.10	1206	H	1.25	1.05	1.49	0.0123	1.25	1.04	1.51	0.010905
ARL14	rs76033371	A/G	3	0.09	3939	All	CM	0.10	1222	H	1.18	1.00	1.40	0.0529	1.19	1.00	1.42	0.0452
B3GALNT1	rs12107243	G/C	3	0.49	3934	All	CM	0.48	1227	D	0.91	0.79	1.06	0.2386	0.93	0.80	1.09	0.23785
IL17RD	rs6780995	A/G	3	0.41	3939	All	CM	0.39	1223	A	0.92	0.84	1.01	0.0763	0.93	0.84	1.02	0.0880
IL17RE	rs708567	A/G	3	0.50	3881	All	CM	0.49	1220	A	0.98	0.90	1.08	0.6937	0.95	0.81	1.10	0.65809
TLR9	rs187084	T/C	3	0.33	3933	All	CM	0.32	1095	A	0.92	0.83	1.02	0.0973	0.89	0.80	0.99	0.0894
FREM3	rs149914432	A/C	4	0.10	3915	All	CM	0.06	1221	A	0.60	0.50	0.73	2.12×10 ⁻⁸	0.64	0.50	0.81	3.08×10 ⁻⁸
FREM3	rs184895969	C/T	4	0.00	3940	All	CM	0.00	1230	H	0.73	0.33	1.60	0.4133	0.82	0.35	1.95	0.4217
FREM3	rs184908374	G/A	4	0.00	3943	All	CM	0.00	1230	H	0.90	0.40	2.01	0.7950	1.02	0.42	2.46	0.80742
FREM3	rs186790584	A/T	4	0.10	3941	All	CM	0.06	1229	A	0.60	0.50	0.72	1.44×10 ⁻⁸	0.61	0.50	0.8	2.38×10 ⁻⁸
FREM3	rs186873296	A/G	4	0.10	3926	All	CM	0.06	1225	A	0.59	0.49	0.71	7.16×10 ⁻⁹	0.63	0.49	0.79	1.12×10 ⁻⁸
GAB1	rs7663712	A/G	4	0.02	3938	All	CM	0.03	1229	A	1.18	0.89	1.57	0.2524	1.15	0.85	1.54	0.2709
INPP4B	rs13103597	C/T	4	0.27	3925	All	CM	0.23	1229	A	0.81	0.73	0.91	0.0002	0.88	0.78	0.98	0.00031243
INPP4B	rs77389579	G/T	4	0.07	3937	All	CM	0.05	1223	A	0.63	0.51	0.78	1.48×10 ⁻⁵	0.66	0.52	0.83	1.64×10 ⁻⁵
TLR1	rs4833095	C/T	4	0.09	3921	All	CM	0.10	1095	R	1.39	0.72	2.70	0.3349	1.35	0.69	2.67	0.33249
TLR6	rs5743809	T/C	4	0.06	3884	All	CM	0.06	1045	H	1.07	0.86	1.33	0.5686	0.73	0.20	2.60	0.6295
TLR6	rs5743810	C/T	4	0.02	3936	All	CM	0.01	1098	A	0.71	0.46	1.09	0.1017	0.62	0.39	0.97	0.10323
USP38	rs28459062	T/C	4	0.16	3931	All	CM	0.18	1230	A	1.16	1.03	1.31	0.0130	1.15	1.02	1.31	0.0169
USP38	rs4266246	C/T	4	0.22	3928	All	CM	0.22	1228	H	1.08	0.94	1.24	0.2589	1.03	0.89	1.19	0.23556
C6	rs1801033	A/C	5	0.47	3927	All	CM	0.48	1093	D	1.07	0.92	1.24	0.3939	1.06	0.91	1.24	0.3683
IL13	rs20541	C/T	5	0.22	3796	All	CM	0.21	1052	R	0.77	0.55	1.08	0.1217	0.75	0.53	1.06	0.14752
IRF	rs2706384	C/A	5	0.38	3819	All	CM	0.35	1072	D	0.90	0.78	1.04	0.1448	0.90	0.78	1.04	0.1709
CTLA	rs2242665	A/G	6	0.26	3853	All	CM	0.26	1077	H	1.02	0.89	1.18	0.7356	0.92	0.69	1.22	0.7311
GABBR1	rs192151845	G/A	6	0.01	3943	All	CM	0.01	1226	H	1.25	0.76	2.08	0.3858	1.31	0.78	2.19	0.3749
HCG4	rs114980857	C/T	6	0.21	3843	All	CM	0.22	1222	D	1.06	0.93	1.21	0.4043	1.04	0.91	1.20	0.40318
HSPA1B	rs6457452	C/T	6	0.19	3917	All	CM	0.18	1211	H	0.90	0.78	1.04	0.1480	0.92	0.79	1.06	0.1841
IL20RA	rs1555498	C/T	6	0.46	3947	All	CM	0.48	1222	R	1.03	0.88	1.20	0.7074	1.03	0.87	1.20	0.74783
LTA	rs2239704	G/T	6	0.12	3896	All	CM	0.13	1206	R	1.36	0.82	2.24	0.2373	1.58	0.94	2.67	0.2342
LTA	rs909253	T/C	6	0.49	3921	All	CM	0.49	1219	H	1.04	0.91	1.18	0.5970	1.06	0.93	1.22	0.5589
PLEKHG1	rs114726617	G/A	6	0.04	3940	All	CM	0.04	1230	A	1.20	0.95	1.50	0.1254	1.26	0.99	1.60	0.1354
PLEKHG1	rs141555199	C/G	6	0.02	3943	All	CM	0.02	1230	R	-	-	-	-	-	-	-	-
PLEKHG1	rs142712208	G/A	6	0.03	3927	All	CM	0.04	1231	D	1.21	0.94	1.57	0.1408	1.29	1.00	1.68	0.1658
PLEKHG1	rs15116938	C/T	6	0.03	3939	All	CM	0.04	1230	A	1.15	0.90	1.46	0.2823	1.23	0.95	1.59	0.32486
PLEKHG1	rs151293197	C/G	6	0.03	3940	All	CM	0.04	1230	A	1.16	0.90	1.48	0.2509	1.24	0.95	1.60	0.2907
PLEKHG1	rs2131263	C/T	6	0.04	3938	All	CM	0.04	1230	A	1.20	0.95	1.50	0.1261	1.26	0.99	1.60	0.13612
PLEKHG1	rs55958968	A/G	6	0.05	3932	All	CM	0.06	1230	H	1.17	0.94	1.46	0.1558	1.12	0.90	1.38	0.2841
PLEKHG1	rs76924464	G/A	6	0.03	3940	All	CM	0.04	1230	A	1.17	0.92	1.49	0.2128	1.25	0.97	1.62	0.24749
PLEKHG1	rs79100774	G/C	6	0.03	3939	All	CM	0.04	1230	A	1.17	0.92	1.49	0.2121	1.25	0.97	1.62	0.2448
SNORD48	rs116288147	C/T	6	0.24	3896	All	CM	0.24	1210	R	1.13	0.87	1.48	0.3661	1.15	0.86	1.53	0.49978
TNF	rs1799964	T/C	6	0.27	3938	All	CM	0.28	1210	R	1.07	0.84	1.36	0.5924	1.09	0.85	1.41	0.5856
TNF	rs1800629	G/A	6	0.09	3940	All	CM	0.09	1225	D	0.97	0.82	1.15	0.7156	0.97	0.82	1.16	0.76027
TNF	rs1800750	G/A	6	0.05	3868	All	CM	0.05	737	D	0.89	0.68	1.17	0.3967	0.90	0.68	1.19	0.3455
TNF	rs3093662	A/G	6	0.11	3936	All	CM	0.12	1219	R	0.70	0.37	1.32	0.2524	0.69	0.36	1.31	0.25352
TNF	rs361525	G/A																

<i>MKI67</i>	rs148494166	T/C	10	0.00	3913	All	CM	0.00	1193	H	1.89	0.82	4.37	0.1512	1.59	0.67	3.74	0.14647
<i>HBB</i>	rs334	A/T	11	0.08	3949	All	CM	0.02	1222	H	0.14	0.09	0.21	8.64×10 ⁻³⁹	0.14	0.09	0.21	1.67×10 ⁻³⁸
<i>RRM1</i>	rs116472045	A/G	11	0.02	3939	All	CM	0.01	1227	A	0.66	0.43	1.03	0.0559	0.60	0.38	0.96	0.046517
<i>RTN3</i>	rs542998	T/C	11	0.43	3929	All	CM	0.42	1085	A	0.96	0.87	1.06	0.4499	0.99	0.82	1.19	0.4858
<i>TRIM5</i>	rs7935564	G/A	11	0.44	3871	All	CM	0.45	1093	R	1.12	0.94	1.33	0.2020	1.00	0.84	1.19	0.19199
<i>CAND1</i>	rs10459266	C/A	12	0.21	3938	All	CM	0.21	1229	A	1.05	0.94	1.18	0.3494	1.04	0.92	1.16	0.3012
<i>CAND1</i>	rs12307123	C/T	12	0.23	3933	All	CM	0.24	1230	D	1.16	1.02	1.33	0.0242	1.17	1.02	1.34	0.016717
<i>CAND1</i>	rs1566830	G/T	12	0.26	3940	All	CM	0.28	1229	A	1.13	1.02	1.26	0.0174	1.12	1.01	1.25	0.0132
<i>GRIP1</i>	rs1394263	T/C	12	0.26	3914	All	CM	0.28	1226	A	1.14	1.03	1.27	0.0116	1.13	1.02	1.26	0.0086331
<i>IL22</i>	rs1012356	A/T	12	0.49	3940	All	CM	0.48	1219	R	0.94	0.80	1.10	0.4187	0.96	0.82	1.13	0.4263
<i>IL22</i>	rs2227478	A/G	12	0.38	3939	All	CM	0.38	1224	R	0.91	0.76	1.09	0.3045	0.88	0.73	1.07	0.3128
<i>IL22</i>	rs2227485	G/A	12	0.42	3928	All	CM	0.41	1219	R	0.92	0.78	1.10	0.3618	0.89	0.74	1.07	0.3404
<i>IL22</i>	rs2227491	C/T	12	0.40	3870	All	CM	0.42	1200	A	1.06	0.96	1.16	0.2375	1.04	0.94	1.15	0.23557
<i>IL22</i>	rs2227507	T/C	12	0.04	3948	All	CM	0.04	1226	H	0.96	0.76	1.22	0.7470	0.97	0.76	1.24	0.7489
<i>TPTE2</i>	rs182873742	C/G	13	0.00	3940	All	CM	0.00	1222	H	3.67	0.22	61.54	0.3809	4.16	0.15	116.27	0.37748
<i>RPS6KLI</i>	rs3742785	C/A	14	0.29	3714	All	CM	0.29	1170	H	1.18	1.03	1.35	0.0143	1.19	1.03	1.37	0.0141
<i>SPTB</i>	rs229587	T/C	14	0.34	3872	All	CM	0.34	1192	R	0.91	0.74	1.12	0.3571	0.91	0.73	1.13	0.40718
<i>YLP1</i>	rs10139016	C/T	14	0.25	3937	All	CM	0.27	1228	D	1.11	0.98	1.27	0.1059	1.10	0.96	1.26	0.1063
<i>ADCY9</i>	rs10775349	C/G	16	0.20	3941	All	CM	0.20	1097	H	1.05	0.91	1.22	0.4706	1.08	0.93	1.25	0.4308
<i>ADCY9</i>	rs2230739	A/G	16	0.09	3944	All	CM	0.07	1098	A	0.84	0.70	1.00	0.0503	0.83	0.69	1.00	0.0479
<i>HBA</i>	α ^{-3.7} -thalassaemia	I/D	16	0.41	3946	All	CM	0.37	1189	A	0.83	0.75	0.91	0.0001	0.83	0.76	0.90	0.0001
<i>ILA</i>	rs2243250	T/C	16	0.20	3902	All	CM	0.22	1212	D	1.13	0.98	1.29	0.0855	1.11	0.96	1.28	0.0858
<i>ILAR</i>	rs1805015	T/C	16	0.42	3927	All	CM	0.44	1223	D	1.12	0.98	1.29	0.1027	1.15	0.99	1.33	0.093058
<i>ADORA2B</i>	rs2535611	T/C	17	0.10	3911	All	CM	0.12	1094	D	1.10	0.93	1.30	0.2651	1.09	0.92	1.29	0.2820
<i>NOS2</i>	rs1800482	G/C	17	0.08	3922	All	CM	0.07	1218	H	0.94	0.78	1.14	0.5488	0.94	0.78	1.13	0.50249
<i>NOS2</i>	rs2297518	G/A	17	0.14	3910	All	CM	0.13	1091	A	0.92	0.80	1.06	0.2360	0.93	0.80	1.08	0.2342
<i>NOS2</i>	rs8078340	C/T	17	0.21	3948	All	CM	0.20	1225	R	0.66	0.45	0.97	0.0275	0.64	0.43	0.94	0.028385
<i>NOS2</i>	rs9282799	C/T	17	0.03	3949	All	CM	0.03	1220	H	1.29	0.98	1.69	0.0735	1.27	0.95	1.70	0.0796
<i>TBX2</i>	rs73991577	A/G	17	0.00	3937	All	CM	0.00	1230	H	0.57	0.19	1.70	0.2890	0.60	0.20	1.80	0.2945
<i>EMR1</i>	rs373533	G/T	19	0.46	3863	All	CM	0.49	1091	D	1.14	0.98	1.33	0.0908	1.12	0.95	1.32	0.1102
<i>EMR1</i>	rs461645	C/T	19	0.47	3933	All	CM	0.49	1097	D	1.14	0.98	1.33	0.0924	1.12	0.95	1.31	0.11209
<i>ICAM</i>	rs5498	A/G	19	0.11	3908	All	CM	0.13	1204	H	1.08	0.92	1.26	0.3494	1.05	0.89	1.24	0.3330
<i>GNAS</i>	rs8386	C/T	20	0.19	3944	All	CM	0.19	1223	R	1.46	1.05	2.03	0.0299	1.42	1.01	1.99	0.029177
<i>DERL3</i>	rs1128127	G/A	22	0.50	3911	All	CM	0.47	1221	R	0.92	0.79	1.08	0.3098	0.91	0.77	1.07	0.3245
<i>CD40LG</i>	rs1126535	T/C	X	0.15	3934	M+F	CM	0.17	1223	R	1.23	0.99	1.54	0.0649	1.20	0.95	1.51	0.089102
<i>CD40LG</i>	rs1126535	T/C	X	0.16	1947	F	CM	0.15	601	D	0.94	0.76	1.15	0.5486	0.95	0.77	1.16	0.5686
<i>CD40LG</i>	rs1126535	C/T	X	0.85	1987	M	CM	0.81	622	HM	0.77	0.61	0.98	0.0348	0.79	0.62	0.99	0.044779
<i>CD40LG</i>	rs3092945	T/C	X	0.21	3927	M+F	CM	0.25	1223	R	1.32	1.10	1.59	0.0034	1.38	1.14	1.67	0.0037
<i>CD40LG</i>	rs3092945	T/C	X	0.21	1942	F	CM	0.25	602	R	1.89	1.30	2.76	0.0012	1.23	1.05	1.43	0.007
<i>CD40LG</i>	rs3092945	C/T	X	0.78	1985	M	CM	0.74	621	HM	0.84	0.68	1.04	0.1097	0.81	0.66	1.00	0.0500
<i>G6PD</i>	rs1050828	C/T	X	0.19	3940	M+F	CM	0.18	1220	H	0.82	0.70	0.97	0.0100	0.80	0.68	0.96	0.0108
<i>G6PD</i>	rs1050828	C/T	X	0.20	1951	F	CM	0.18	600	H	0.80	0.65	0.98	0.0202	0.79	0.64	0.97	0.0200
<i>G6PD</i>	rs1050828	C/T	X	0.19	1989	M	CM	0.18	620	HM	0.92	0.72	1.17	0.4891	0.91	0.72	1.15	0.45
<i>G6PD</i>	rs1050829	T/C	X	0.40	3928	M+F	CM	0.41	1221	H	0.84	0.70	1.01	0.0609	0.87	0.71	1.06	0.0658
<i>G6PD</i>	rs1050829	T/C	X	0.41	1952	F	CM	0.41	601	H	0.84	0.70	1.01	0.0631	0.85	0.72	1.10	0.083127
<i>G6PD</i>	rs1050829	C/T	X	0.60	1976	M	CM	0.58	620	HM	0.94	0.78	1.14	0.5353	0.95	0.79	1.17	0.5353

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and rs334 genotype. All individuals having cerebral malaria were included in this analysis even if they were identified with multiple severe phenotypes.

Table S4: Odds ratios for all-CM for all polymorphisms investigated.

Gene	SNPID	Alleles (ancestral / derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	MO	Unadjusted				Adjusted [§]			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.08	3940	All	SMA	0.07	666	H	0.85	0.66	1.10	0.2075	0.82	0.63	1.07	0.2054
AJAP1	rs6674631	G/A	1	0.13	3928	All	SMA	0.11	667	A	0.84	0.70	1.00	0.0502	0.82	0.67	0.99	0.045017
ATP2B4	rs10900585	G/T	1	0.34	3762	All	SMA	0.31	659	R	1.52	1.11	2.09	0.0034	1.00	0.54	1.85	0.0040
ATP2B4	rs1541255	A/G	1	0.33	3933	All	SMA	0.29	665	R	0.63	0.46	0.87	0.0030	0.61	0.43	0.84	0.003432
ATP2B4	rs3753036	G/A	1	0.04	3939	All	SMA	0.05	667	D	1.21	0.91	1.60	0.1976	1.19	0.88	1.61	0.2236
ATP2B4	rs4951074	G/A	1	0.32	3934	All	SMA	0.29	666	R	0.66	0.48	0.91	0.0087	0.67	0.53	0.92	0.009973
ATP2B4	rs55868763	G/C	1	0.33	3931	All	SMA	0.29	661	R	0.62	0.45	0.85	0.0019	0.64	0.48	0.89	0.0021
CR1	rs17047660	A/G	1	0.16	3923	All	SMA	0.15	683	H	0.87	0.72	1.05	0.1335	0.85	0.69	1.03	0.096844
CR1	rs17047661	G/A	1	0.32	3934	All	SMA	0.33	685	A	1.06	0.93	1.20	0.3728	1.06	0.93	1.21	0.3946
DARC	rs2814778	G/A	1	0.00	3944	All	SMA	0.00	556	H	0.76	0.17	3.41	0.7084	0.74	0.16	3.35	0.71045
GBP7	rs1803632	G/C	1	0.49	3942	All	SMA	0.50	685	R	1.10	0.91	1.33	0.3224	1.10	0.90	1.34	0.4239
IL10	rs1800890	A/T	1	0.24	3945	All	SMA	0.24	687	R	0.83	0.58	1.19	0.3086	0.89	0.62	1.30	0.3312
IL10	rs1800896	T/C	1	0.38	3943	All	SMA	0.38	682	R	1.05	0.84	1.33	0.6612	1.08	0.84	1.37	0.5771
IL10	rs3024500	G/A	1	0.45	3928	All	SMA	0.47	682	H	0.80	0.68	0.95	0.0094	0.81	0.68	0.97	0.012456
LPHN2	rs146428334	A/G	1	0.00	3930	All	SMA	0.00	668	H	0.60	0.18	1.99	0.3751	0.58	0.17	1.93	0.3811
LPHN2	rs4650365	T/C	1	0.20	3936	All	SMA	0.18	667	D	0.85	0.71	1.01	0.0586	0.86	0.72	1.04	0.056572
LPHN2	rs72933304	C/A	1	0.09	3942	All	SMA	0.07	666	A	0.85	0.68	1.06	0.1517	0.88	0.70	1.11	0.1714
LPHN2	rs72933310	T/C	1	0.09	3927	All	SMA	0.08	664	A	0.90	0.73	1.12	0.3428	0.92	0.74	1.14	0.33259
LPHN2	rs72933350	T/C	1	0.05	3940	All	SMA	0.04	667	D	0.72	0.52	0.98	0.0303	0.79	0.57	1.10	0.0344
IL1A	rs17561	G/T	2	0.16	3938	All	SMA	0.17	687	R	1.05	0.63	1.76	0.8487	1.06	0.61	1.83	0.82324
IL1B	rs1143634	C/T	2	0.11	3941	All	SMA	0.13	686	A	1.10	0.92	1.30	0.3102	1.06	0.88	1.28	0.3764
LAPTM4A	rs973128	C/G	2	0.44	3885	All	SMA	0.46	658	A	1.11	0.98	1.25	0.0888	1.11	0.98	1.26	0.079476
LOC727982	rs10188961	A/G	2	0.40	3934	All	SMA	0.45	665	D	1.44	1.20	1.72	0.0001	1.48	1.22	1.80	0.00005
LOC727982	rs1371474	T/C	2	0.27	3933	All	SMA	0.30	667	H	1.32	1.11	1.55	0.0013	1.42	1.19	1.69	0.001172
LOC727982	rs1371478	C/T	2	0.27	3909	All	SMA	0.30	663	D	1.31	1.11	1.55	0.0014	1.45	1.22	1.74	0.0011
SDCI	rs11899121	C/G	2	0.47	3921	All	SMA	0.47	666	D	0.97	0.81	1.16	0.7102	1.01	0.90	1.15	0.91039
ZSWIM2	rs144778284	C/T	2	0.00	3937	All	SMA	0.00	667	R	-	-	-	-	-	-	-	-
ZSWIM2	rs4316902	A/G	2	0.18	3910	All	SMA	0.18	666	R	0.87	0.53	1.42	0.5709	0.88	0.52	1.48	0.58979
ARL14	rs74954675	A/C	3	0.08	3907	All	SMA	0.10	662	H	1.31	1.05	1.62	0.0192	1.27	1.00	1.60	0.0243
ARL14	rs75731597	A/C	3	0.08	3491	All	SMA	0.10	646	H	1.27	1.02	1.59	0.0353	1.30	1.07	1.59	0.046063
ARL14	rs76033371	A/G	3	0.09	3939	All	SMA	0.11	661	H	1.26	1.02	1.56	0.0353	1.20	0.95	1.50	0.0431
B3GALNT1	rs12107243	G/C	3	0.49	3934	All	SMA	0.49	667	D	1.02	0.84	1.23	0.8608	1.03	0.91	1.17	0.92778
ILI7RD	rs6780995	A/G	3	0.41	3939	All	SMA	0.39	687	R	0.84	0.67	1.05	0.1222	0.87	0.68	1.10	0.1597
ILI7RE	rs708567	A/G	3	0.50	3881	All	SMA	0.49	685	D	0.95	0.79	1.15	0.5998	0.94	0.77	1.15	0.54384
TLR9	rs187084	T/C	3	0.33	3933	All	SMA	0.33	557	H	0.95	0.79	1.14	0.5787	0.95	0.79	1.15	0.5583
FREM3	rs149914432	A/C	4	0.10	3915	All	SMA	0.06	663	A	0.58	0.45	0.74	2.26x10 ⁻⁶	0.68	0.50	0.94	3.68x10 ⁻⁶
FREM3	rs184895969	C/T	4	0.00	3940	All	SMA	0.00	668	H	1.02	0.42	2.47	0.9662	1.26	0.48	3.27	0.9530
FREM3	rs184908374	G/A	4	0.00	3943	All	SMA	0.00	666	H	1.25	0.51	3.07	0.6373	1.62	0.61	4.31	0.62615
FREM3	rs186790584	A/T	4	0.10	3941	All	SMA	0.06	667	A	0.57	0.44	0.73	1.55x10 ⁻⁶	0.63	0.48	0.8	2.72x10 ⁻⁶
FREM3	rs186873296	A/G	4	0.10	3926	All	SMA	0.05	663	A	0.55	0.43	0.70	4.56x10 ⁻⁷	0.67	0.44	0.83	8.20x10 ⁻⁷
GAB1	rs7663712	A/G	4	0.02	3938	All	SMA	0.03	668	R	-	-	-	-	1.25	0.85	1.82	0.3793
INPP4B	rs13103597	C/T	4	0.27	3925	All	SMA	0.22	666	D	0.76	0.64	0.90	0.0013	0.75	0.64	0.89	0.001257
INPP4B	rs77389579	G/T	4	0.07	3937	All	SMA	0.04	666	A	0.56	0.42	0.75	3.57x10 ⁻⁵	0.60	0.43	0.82	5.35x10 ⁻⁵
TLR1	rs4833095	C/T	4	0.09	3921	All	SMA	0.08	558	H	0.79	0.60	1.03	0.0697	0.84	0.66	1.07	0.44251
TLR6	rs5743809	T/C	4	0.06	3884	All	SMA	0.05	533	H	0.89	0.65	1.20	0.4320	0.89	0.64	1.24	0.3248
TLR6	rs5743810	C/T	4	0.02	3936	All	SMA	0.01	558	A	0.78	0.44	1.37	0.3732	0.72	0.40	1.28	0.37936
USP38	rs28459062	T/C	4	0.16	3931	All	SMA	0.17	666	A	1.07	0.92	1.25	0.3788	1.04	0.88	1.22	0.4579
USP38	rs4266246	C/T	4	0.22	3928	All	SMA	0.23	667	R	1.23	0.86	1.77	0.2740	1.14	0.78	1.66	0.25133
C6	rs1801033	A/C	5	0.47	3927	All	SMA	0.46	556	R	0.92	0.74	1.14	0.4372	0.89	0.71	1.12	0.4362
IL13	rs20541	C/T	5	0.22	3796	All	SMA	0.22	535	R	0.63	0.38	1.03	0.0520	0.59	0.35	1.00	0.062928
IRF	rs2706384	C/A	5	0.38	3819	All	SMA	0.37	546	H	1.25	1.04	1.49	0.0164	1.28	1.06	1.55	0.0102
CTLA	rs2242665	A/G	6	0.26	3853	All	SMA	0.25	550	H	0.86	0.71	1.03	0.1016	0.81	0.67	0.99	0.080986
GABBR1	rs192151845	G/A	6	0.01	3943	All	SMA	0.01	665	H	1.13	0.59	2.18	0.7198	1.15	0.59	2.25	0.7063
HCG4	rs114980857	C/T	6	0.21	3843	All	SMA	0.21	654	R	0.87	0.57	1.31	0.4867	0.81	0.52	1.26	0.50211
HSPA1B	rs6457452	C/T	6	0.19	3917	All	SMA	0.17	674	D	0.84	0.70	1.00	0.0478	0.81	0.66	0.99	0.0443
IL20RA	rs1555498	C/T	6	0.46	3947	All	SMA	0.50	684	R	1.16	0.96	1.40	0.1235	1.17	0.96	1.43	0.1376
LTA	rs2239704	G/T	6	0.12	3896	All	SMA	0.11	670	H	0.86	0.70	1.06	0.1431	0.81	0.65	1.01	0.1256
LTA	rs909253	T/C	6	0.49	3921	All	SMA	0.50	680	D	1.10	0.91	1.33	0.3397	1.22	0.99	1.49	0.25607
PLEKHG1	rs114726617	G/A	6	0.04	3940	All	SMA	0.04	668	R	2.32	0.60	8.93	0.2513	2.99	0.74	12.04	0.2482
PLEKHG1	rs141555199	C/G	6	0.02	3943	All	SMA	0.02	667	R	-	-	-	-	-	-	-	-
PLEKHG1	rs142712208	G/A	6	0.03	3927	All	SMA	0.04	667	R	3.51	0.87	14.27	0.1071	5.40	1.28	22.73	0.1055
PLEKHG1	rs15116938	C/T	6	0.03	3939	All	SMA	0.04	668	R	4.83	1.34	17.36	0.0270	7.43	1.99	27.71	0.026517
PLEKHG1	rs151293197	C/G	6	0.03	3940	All	SMA	0.04	667	R	3.52	0.87	14.30	0.1065	5.40	1.28	22.76	0.1049
PLEKHG1	rs2131263	C/T	6	0.04	3938	All	SMA	0.04	668	R	2.32	0.60	8.92	0.2515	2.99	0.74	12.04	0.24833
PLEKHG1	rs55958968	A/G	6	0.05	3932	All	SMA	0.06	668	H	1.21	0.93	1.58	0.1617	1.27	0.96	1.67	0.1399
PLEKHG1	rs76924464	G/A	6	0.03	3940	All	SMA	0.04	668	R	3.51	0.87	14.26	0.1071	5.37	1.28	22.62	0.10551
PLEKHG1	rs79100774	G/C	6	0.03	3939	All	SMA	0.04	665	R	3.53	0.87	14.32	0.1060	5.39	1.28	22.72	0.1044
SNORD48	rs116288147	C/T	6	0.24	3896	All	SMA	0.22	658	D	0.89	0.75	1.06	0.1831	0.86	0.72	1.03	0.17358
TNF	rs1799964	T/C	6	0.27	3938	All	SMA	0.28	674	R	1.10	0.81	1.49	0.5481	1.07	0.78	1.48	0.5174
TNF	rs1800629	G/A	6	0.09	3940	All	SMA	0.10	685	R	1.92	0.90	4.09	0.1097	1.84	0.81	4.15	0.19385
TNF	rs1800750	G/A	6	0.05	3868	All	SMA	0.05	423	R	0.53	0.07	4.01	0.5007	0.72	0.09	5.54	0.5090
TNF	rs3093662	A/G	6	0.11	3936	All	SMA	0.10	682	R	0.43	0.16	1.20	0.0698	0.45	0.16	1.26	0.072694
TNF	rs361525	G/A	6	0.07	3948	All	SMA	0.06	684	R	-	-	-	-	-	-	-	-

<i>MKI67</i>	rs148494166	T/C	10	0.00	3913	All	SMA	0.00	645	H	0.40	0.05	3.04	0.3102	0.33	0.04	2.54	0.31262
<i>HBB</i>	rs334	A/T	11	0.08	3949	All	SMA	0.02	683	H	0.07	0.04	0.15	2.11×10 ⁻³¹	0.07	0.03	0.17	2.61×10 ⁻³¹
<i>RRM1</i>	rs116472045	A/G	11	0.02	3939	All	SMA	0.02	666	R	-	-	-	-	-	-	-	-
<i>RTN3</i>	rs542998	T/C	11	0.43	3929	All	SMA	0.45	546	D	1.14	0.94	1.39	0.1843	1.15	0.94	1.41	0.1740
<i>TRIM5</i>	rs7935564	G/A	11	0.44	3871	All	SMA	0.47	554	D	1.24	1.01	1.51	0.0381	1.18	0.95	1.47	0.032002
<i>CAND1</i>	rs10459266	C/A	12	0.21	3938	All	SMA	0.23	667	D	1.22	1.03	1.44	0.0218	1.23	1.03	1.48	0.0210
<i>CAND1</i>	rs12307123	C/T	12	0.23	3933	All	SMA	0.24	666	D	1.14	0.97	1.35	0.1207	1.16	0.97	1.38	0.083643
<i>CAND1</i>	rs1566830	G/T	12	0.26	3940	All	SMA	0.28	666	A	1.13	0.99	1.29	0.0768	1.13	0.98	1.30	0.0594
<i>GRIP1</i>	rs1394263	T/C	12	0.26	3914	All	SMA	0.28	661	A	1.13	0.99	1.29	0.0741	1.13	0.99	1.30	0.057273
<i>IL22</i>	rs1012356	A/T	12	0.49	3940	All	SMA	0.49	685	H	1.18	1.00	1.39	0.0455	1.17	0.99	1.40	0.0560
<i>IL22</i>	rs2227478	A/G	12	0.38	3939	All	SMA	0.38	686	R	0.85	0.67	1.08	0.1800	0.87	0.68	1.12	0.20618
<i>IL22</i>	rs2227485	G/A	12	0.42	3928	All	SMA	0.41	681	H	1.27	1.08	1.50	0.0038	1.27	1.07	1.51	0.0060
<i>IL22</i>	rs2227491	C/T	12	0.40	3870	All	SMA	0.40	676	H	1.13	0.96	1.33	0.1469	0.82	0.64	1.06	0.17937
<i>IL22</i>	rs2227507	T/C	12	0.04	3948	All	SMA	0.04	687	A	0.83	0.61	1.12	0.2136	0.84	0.61	1.16	0.2545
<i>TPTE2</i>	rs182873742	C/G	13	0.00	3940	All	SMA	0.00	663	H	6.43	0.39	104.8	0.2186	5.95	0.26	135.3	0.21702
<i>RPS6KL1</i>	rs3742785	C/A	14	0.29	3714	All	SMA	0.32	650	D	1.33	1.12	1.57	0.0010	1.37	1.15	1.64	0.0010
<i>SPTB</i>	rs229587	T/C	14	0.34	3872	All	SMA	0.35	660	R	1.13	0.88	1.45	0.3452	1.07	0.81	1.39	0.33503
<i>YLPM1</i>	rs10139016	C/T	14	0.25	3937	All	SMA	0.28	666	D	1.25	1.06	1.47	0.0095	1.26	1.06	1.51	0.0104
<i>ADCY9</i>	rs10775349	C/G	16	0.20	3941	All	SMA	0.21	557	H	1.08	0.89	1.30	0.4300	1.06	0.87	1.30	0.40226
<i>ADCY9</i>	rs2230739	A/G	16	0.09	3944	All	SMA	0.08	558	R	0.49	0.12	2.07	0.2842	0.50	0.12	2.13	0.2861
<i>HBA</i>	α ^{-3.7} -thalassaemia	I/D	16	0.41	3946	All	SMA	0.34	639	A	0.72	0.64	0.82	1.55×10 ⁻⁶	0.72	0.65	0.84	5.45×10 ⁻⁶
<i>IL4</i>	rs2243250	T/C	16	0.20	3902	All	SMA	0.21	677	R	0.69	0.43	1.11	0.1067	1.02	0.87	1.19	0.6437
<i>IL4R</i>	rs1805015	T/C	16	0.42	3927	All	SMA	0.42	684	D	0.97	0.82	1.15	0.7331	1.03	0.86	1.24	0.64537
<i>ADORA2B</i>	rs2535611	T/C	17	0.10	3911	All	SMA	0.11	554	A	1.07	0.87	1.30	0.5378	1.01	0.81	1.25	0.5881
<i>NOS2</i>	rs1800482	G/C	17	0.08	3922	All	SMA	0.07	683	R	0.46	0.11	1.96	0.2429	0.22	0.04	1.40	0.24753
<i>NOS2</i>	rs2297518	G/A	17	0.14	3910	All	SMA	0.14	553	R	0.87	0.45	1.71	0.6893	0.98	0.50	1.93	0.7002
<i>NOS2</i>	rs8078340	C/T	17	0.21	3948	All	SMA	0.20	686	A	0.96	0.83	1.11	0.5666	0.93	0.80	1.08	0.42929
<i>NOS2</i>	rs9282799	C/T	17	0.03	3949	All	SMA	0.03	685	R	-	-	-	-	-	-	-	-
<i>TBX2</i>	rs73991577	A/G	17	0.00	3937	All	SMA	0.00	667	H	0.27	0.04	2.03	0.1199	0.33	0.04	2.54	0.12192
<i>EMR1</i>	rs373533	G/T	19	0.46	3863	All	SMA	0.49	551	D	1.14	0.93	1.40	0.2161	1.10	0.89	1.36	0.2489
<i>EMR1</i>	rs461645	C/T	19	0.47	3933	All	SMA	0.49	558	A	1.07	0.95	1.22	0.2667	1.06	0.93	1.21	0.24864
<i>ICAM</i>	rs5498	A/G	19	0.11	3908	All	SMA	0.12	677	H	1.07	0.88	1.31	0.4818	1.02	0.83	1.27	0.5481
<i>GNAS</i>	rs8386	C/T	20	0.19	3944	All	SMA	0.19	686	R	1.39	0.92	2.09	0.1278	1.35	0.88	2.07	0.11859
<i>DERL3</i>	rs1128127	G/A	22	0.50	3911	All	SMA	0.47	684	R	0.88	0.72	1.07	0.1880	0.84	0.68	1.04	0.2062
<i>CD40LG</i>	rs1126535	T/C	X	0.15	3934	M+F	SMA	0.16	686	H	0.75	0.56	0.99	0.0416	0.75	0.56	1.02	0.071258
<i>CD40LG</i>	rs1126535	T/C	X	0.16	1947	F	SMA	0.13	328	D	0.75	0.57	0.99	0.0393	0.77	0.61	0.99	0.0393
<i>CD40LG</i>	rs1126535	C/T	X	0.85	1987	M	SMA	0.81	358	HM	0.74	0.55	1.00	0.0528	0.76	0.58	1.10	0.056789
<i>CD40LG</i>	rs3092945	T/C	X	0.21	3927	M+F	SMA	0.26	683	A	1.17	1.05	1.31	0.0038	1.18	1.06	1.33	0.0031
<i>CD40LG</i>	rs3092945	T/C	X	0.21	1942	F	SMA	0.26	328	A	1.33	1.10	1.62	0.0037	1.36	1.12	1.64	0.001
<i>CD40LG</i>	rs3092945	C/T	X	0.78	1985	M	SMA	0.74	355	HM	0.81	0.63	1.05	0.1224	0.79	0.61	1.02	0.0700
<i>G6PD</i>	rs1050828	C/T	X	0.19	3940	M+F	SMA	0.23	680	R	1.60	1.26	2.03	0.0002	1.73	1.34	2.23	0.000171
<i>G6PD</i>	rs1050828	C/T	X	0.20	1951	F	SMA	0.17	325	H	0.79	0.61	1.03	0.0743	0.78	0.60	1.02	0.0600
<i>G6PD</i>	rs1050828	C/T	X	0.19	1989	M	SMA	0.28	355	HM	1.72	1.32	2.23	6.98×10 ⁻⁵	1.68	1.30	2.17	0.0001
<i>G6PD</i>	rs1050829	T/C	X	0.40	3928	M+F	SMA	0.45	680	R	1.30	1.08	1.56	0.0054	1.16	1.03	1.31	0.0138
<i>G6PD</i>	rs1050829	T/C	X	0.41	1952	F	SMA	0.41	325	R	1.06	0.77	1.45	0.7268	1.09	0.79	1.55	0.72678
<i>G6PD</i>	rs1050829	C/T	X	0.60	1976	M	SMA	0.51	355	HM	0.69	0.55	0.87	0.0016	0.71	0.58	0.90	0.0017

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and rs334 genotype. All individuals having severe malarial anaemia were included in this analysis even if they were identified with multiple severe phenotypes.

Table S5: Odds ratios for all SMA for all polymorphisms investigated.

Gene	SNPID	Alleles (ancestral / derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	MO	Unadjusted				Adjusted ⁸			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.08	3940	All	RD	0.06	683	A	0.81	0.64	1.02	0.0701	0.71	0.43	1.16	0.1850
AJAP1	rs6674631	G/A	1	0.13	3928	All	RD	0.12	683	A	0.90	0.75	1.07	0.2318	1.33	0.92	1.93	0.15938
ATP2B4	rs10900585	G/T	1	0.34	3762	All	RD	0.32	680	R	1.38	1.02	1.87	0.0155	1.13	0.52	2.44	0.0151
ATP2B4	rs1541255	A/G	1	0.33	3933	All	RD	0.30	684	R	0.62	0.46	0.85	0.0019	0.64	0.46	0.88	0.002
ATP2B4	rs3753036	G/A	1	0.04	3939	All	RD	0.04	684	H	0.82	0.59	1.13	0.2162	0.83	0.60	1.14	0.2218
ATP2B4	rs4951074	G/A	1	0.32	3934	All	RD	0.29	683	R	0.65	0.47	0.90	0.0059	0.66	0.52	0.91	0.0057
ATP2B4	rs55868763	G/C	1	0.33	3931	All	RD	0.30	683	R	0.63	0.47	0.87	0.0025	0.65	0.54	0.90	0.0030
CR1	rs17047660	A/G	1	0.16	3923	All	RD	0.17	683	A	1.10	0.94	1.29	0.2219	1.22	0.90	1.66	0.33548
CR1	rs17047661	G/A	1	0.32	3934	All	RD	0.34	683	A	1.08	0.95	1.22	0.2427	1.22	0.73	2.03	0.2094
DARC	rs2814778	G/A	1	0.00	3944	All	RD	0.00	601	H	0.83	0.18	3.71	0.8003	0.87	0.22	4.21	0.34691
GBP7	rs1803632	G/C	1	0.49	3942	All	RD	0.52	683	R	1.22	1.01	1.47	0.0361	1.23	1.10	1.53	0.0360
IL10	rs1800890	A/T	1	0.24	3945	All	RD	0.23	685	R	0.75	0.51	1.09	0.1200	0.75	0.51	1.11	0.12
IL10	rs1800896	T/C	1	0.38	3943	All	RD	0.35	684	D	0.89	0.76	1.05	0.1787	0.74	0.44	1.25	0.2693
IL10	rs3024500	G/A	1	0.45	3928	All	RD	0.48	682	A	1.12	0.99	1.26	0.0658	1.36	0.98	1.90	0.057556
LPHN2	rs146428334	A/G	1	0.00	3930	All	RD	0.00	684	H	1.12	0.47	2.70	0.8015	1.54	0.36	6.61	0.5752
LPHN2	rs4650365	T/C	1	0.20	3936	All	RD	0.19	684	H	0.88	0.74	1.05	0.1587	0.77	0.28	2.11	0.48287
LPHN2	rs72933304	C/A	1	0.09	3942	All	RD	0.07	683	H	0.83	0.66	1.03	0.0975	0.85	0.68	1.07	0.1122
LPHN2	rs72933310	T/C	1	0.09	3927	All	RD	0.08	683	H	0.89	0.72	1.10	0.2667	1.14	0.75	1.74	0.52604
LPHN2	rs72933350	T/C	1	0.05	3940	All	RD	0.05	684	H	2.35	0.60	9.19	0.2479	1.47	0.92	2.35	0.1538
IL1A	rs17561	G/T	2	0.16	3938	All	RD	0.19	685	A	1.20	1.03	1.39	0.0203	2.39	1.13	5.09	0.064612
IL1B	rs1143634	C/T	2	0.11	3941	All	RD	0.11	683	R	0.78	0.35	1.73	0.5252	0.81	0.55	1.20	0.3628
LAPTM4A	rs973128	C/G	2	0.44	3885	All	RD	0.44	675	H	1.05	0.89	1.24	0.5620	1.07	0.90	1.27	0.067
LOC727982	rs10188961	A/G	2	0.40	3934	All	RD	0.43	682	A	1.16	1.03	1.30	0.0165	1.17	0.93	1.47	0.1616
LOC727982	rs1371474	T/C	2	0.27	3933	All	RD	0.29	685	H	1.38	1.17	1.63	0.0001	1.69	1.22	2.35	0.002967
LOC727982	rs1371478	C/T	2	0.27	3909	All	RD	0.29	682	H	1.39	1.18	1.64	0.0001	1.47	1.23	1.74	0.0002
SDC1	rs11899121	C/G	2	0.47	3921	All	RD	0.48	684	H	1.07	0.91	1.26	0.3948	1.27	0.87	1.85	0.29507
ZSWIM2	rs144778284	C/T	2	0.00	3937	All	RD	0.00	684	R	-	-	-	-	-	-	-	-
ZSWIM2	rs4316902	A/G	2	0.18	3910	All	RD	0.16	677	R	0.70	0.41	1.19	0.1706	0.79	0.53	1.16	0.16013
ARL14	rs74954675	A/C	3	0.08	3907	All	RD	0.09	677	H	1.15	0.92	1.44	0.2251	1.05	0.70	1.60	0.8342
ARL14	rs75731597	A/C	3	0.08	3491	All	RD	0.10	670	H	1.23	0.98	1.53	0.0663	1.28	0.99	1.59	0.089
ARL14	rs76033371	A/G	3	0.09	3939	All	RD	0.10	679	H	1.19	0.96	1.47	0.1146	1.08	0.73	1.60	0.7454
B3GALNT1	rs12107243	G/C	3	0.49	3934	All	RD	0.47	683	D	0.87	0.72	1.04	0.1342	0.93	0.74	1.18	0.39756
IL17RD	rs6780995	A/G	3	0.41	3939	All	RD	0.41	685	D	0.98	0.83	1.17	0.8498	0.98	0.63	1.51	0.8753
IL17RE	rs708567	A/G	3	0.50	3881	All	RD	0.49	680	D	0.94	0.78	1.13	0.5045	0.81	0.64	1.02	0.11682
TLR9	rs187084	T/C	3	0.33	3933	All	RD	0.32	601	D	0.87	0.74	1.04	0.1273	0.90	0.81	1.58	0.4593
FREM3	rs149914432	A/C	4	0.10	3915	All	RD	0.06	682	D	0.55	0.43	0.71	9.65×10 ⁻⁷	0.48	0.24	0.95	0.002561
FREM3	rs184895969	C/T	4	0.00	3940	All	RD	0.00	684	H	1.03	0.43	2.48	0.9486	1.04	0.55	3.40	0.0633
FREM3	rs184908374	G/A	4	0.00	3943	All	RD	0.00	684	H	1.05	0.40	2.77	0.9185	1.06	0.50	2.80	0.12848
FREM3	rs186790584	A/T	4	0.10	3941	All	RD	0.06	684	D	0.54	0.42	0.70	6.55×10 ⁻⁷	0.53	0.40	0.7	0.0033
FREM3	rs186873296	A/G	4	0.10	3926	All	RD	0.05	682	D	0.53	0.41	0.69	4.67×10 ⁻⁷	0.61	0.44	0.83	6.45×10 ⁻⁵
GAB1	rs7663712	A/G	4	0.02	3938	All	RD	0.03	684	H	1.21	0.85	1.73	0.3060	1.22	0.19	1.38	0.1530
INPP4B	rs13103597	C/T	4	0.27	3925	All	RD	0.23	683	A	0.78	0.66	0.93	0.0040	0.82	0.72	0.94	0.004952
INPP4B	rs77389579	G/T	4	0.07	3937	All	RD	0.04	683	A	0.58	0.44	0.78	8.53×10 ⁻⁵	0.62	0.46	0.83	9.54×10 ⁻⁵
TLR1	rs4833095	C/T	4	0.09	3921	All	RD	0.10	600	R	1.79	0.84	3.79	0.1499	1.31	0.90	1.89	0.070727
TLR6	rs5743809	T/C	4	0.06	3884	All	RD	0.06	576	R	1.85	0.61	5.67	0.3081	3.71	0.81	17.04	0.1424
TLR6	rs5743810	C/T	4	0.02	3936	All	RD	0.02	602	R	-	-	-	-	-	-	-	-
USP38	rs28459062	T/C	4	0.16	3931	All	RD	0.17	684	R	1.35	0.87	2.11	0.1952	1.05	0.78	1.43	0.5949
USP38	rs4266246	C/T	4	0.22	3928	All	RD	0.22	684	A	1.02	0.88	1.17	0.8077	1.34	0.69	2.61	0.29847
C6	rs1801033	A/C	5	0.47	3927	All	RD	0.48	600	H	1.08	0.91	1.28	0.3912	0.72	0.51	1.02	0.0586
IL13	rs20541	C/T	5	0.22	3796	All	RD	0.22	579	H	0.98	0.81	1.18	0.8427	0.99	0.86	1.48	0.34347
IRF	rs2706384	C/A	5	0.38	3819	All	RD	0.36	586	A	0.94	0.83	1.07	0.3591	1.06	0.67	1.69	0.5697
CTLA	rs2242665	A/G	6	0.26	3853	All	RD	0.28	591	R	1.20	0.87	1.65	0.2804	1.32	1.02	1.70	0.025206
GABBR1	rs192151845	G/A	6	0.01	3943	All	RD	0.01	683	H	1.30	0.70	2.39	0.4154	1.32	0.73	3.67	0.8750
HCG4	rs114980857	C/T	6	0.21	3843	All	RD	0.20	669	A	0.90	0.78	1.04	0.1628	0.89	0.67	1.19	0.57831
HSPA1B	rs6457452	C/T	6	0.19	3917	All	RD	0.19	675	A	0.96	0.83	1.12	0.6348	0.85	0.63	1.16	0.2810
IL20RA	rs1555498	C/T	6	0.46	3947	All	RD	0.48	683	H	0.86	0.73	1.01	0.0613	0.87	0.61	1.24	0.30236
LTA	rs2239704	G/T	6	0.12	3896	All	RD	0.13	673	R	1.60	0.89	2.85	0.1316	1.32	0.91	1.91	0.0873
LTA	rs909253	T/C	6	0.49	3921	All	RD	0.49	680	H	1.13	0.96	1.33	0.1358	1.00	0.79	1.25	0.91093
PLEKHG1	rs114726617	G/A	6	0.04	3940	All	RD	0.05	684	D	1.29	0.96	1.72	0.0980	1.37	0.77	2.44	0.3355
PLEKHG1	rs141555199	C/G	6	0.02	3943	All	RD	0.03	684	H	1.30	0.89	1.89	0.1833	1.32	0.90	1.95	0.67514
PLEKHG1	rs142712208	G/A	6	0.03	3927	All	RD	0.04	685	H	1.36	1.00	1.86	0.0563	1.38	0.74	2.57	0.3974
PLEKHG1	rs15116938	C/T	6	0.03	3939	All	RD	0.04	684	H	1.26	0.92	1.72	0.1537	1.31	0.94	1.82	0.49189
PLEKHG1	rs151293197	C/G	6	0.03	3940	All	RD	0.04	683	H	1.27	0.93	1.73	0.1466	1.32	0.71	2.46	0.4501
PLEKHG1	rs2131263	C/T	6	0.04	3938	All	RD	0.05	684	D	1.29	0.96	1.72	0.0985	1.37	0.77	2.44	0.33629
PLEKHG1	rs55958968	A/G	6	0.05	3932	All	RD	0.06	684	H	1.17	0.90	1.52	0.2575	1.39	0.85	2.25	0.1400
PLEKHG1	rs76924464	G/A	6	0.03	3940	All	RD	0.04	684	H	1.28	0.94	1.74	0.1315	1.29	0.69	2.40	0.48015
PLEKHG1	rs79100774	G/C	6	0.03	3939	All	RD	0.04	683	H	1.25	0.92	1.71	0.1646	1.29	0.69	2.41	0.4816
SNORD48	rs116288147	C/T	6	0.24	3896	All	RD	0.24	675	R	1.05	0.75	1.49	0.7679	1.07	0.81	1.59	0.7787
TNF	rs1799964	T/C	6	0.27	3938	All	RD	0.27	676	R	0.91	0.66	1.26	0.5853	0.95	0.70	1.34	0.6790
TNF	rs1800629	G/A	6	0.09	3940	All	RD	0.09	684	R	1.92	0.90	4.09	0.1094	1.97	0.98	4.95	0.1199
TNF	rs1800750	G/A	6	0.05	3868	All	RD	0.04	352	H	0.71	0.47	1.08	0.0987	0.67	0.25	1.81	0.3678
TNF	rs3093662	A/G	6	0.11	3936	All	RD	0.11	682	R	0.53	0.21	1.33	0.1384	0.87	0.57	1.32	0.54979
TNF	rs361525	G/A	6	0.07	3948	All	RD	0.07	685	R	0.52	0.12	2.22	0.3342	0.54	0.19	3.21	

<i>HBB</i>	rs334	A/T	11	0.08	3949	All	RD	0.02	684	H	0.11	0.06	0.20	8.41×10 ⁻²⁸	0.11	0.06	0.20	1.62×10 ⁻²⁷
<i>RRM1</i>	rs116472045	A/G	11	0.02	3939	All	RD	0.01	682	R	-	-	-	-	-	-	-	-
<i>RTN3</i>	rs542998	T/C	11	0.43	3929	All	RD	0.44	592	R	1.16	0.93	1.43	0.1864	0.82	0.58	1.15	0.3977
<i>TRIM5</i>	rs7935564	G/A	11	0.44	3871	All	RD	0.47	598	A	1.15	1.01	1.30	0.0289	1.29	0.87	1.90	0.091014
<i>CAND1</i>	rs10459266	C/A	12	0.21	3938	All	RD	0.23	684	D	1.22	1.03	1.44	0.0217	1.75	1.26	2.42	0.0009
<i>CAND1</i>	rs12307123	C/T	12	0.23	3933	All	RD	0.25	682	D	1.24	1.05	1.46	0.0113	1.17	0.90	1.52	0.17144
<i>CAND1</i>	rs1566830	G/T	12	0.26	3940	All	RD	0.29	684	A	1.20	1.06	1.37	0.0049	1.76	1.04	2.98	0.0409
<i>GRIP1</i>	rs1394263	T/C	12	0.19	3944	All	RD	0.18	683	H	0.92	0.77	1.09	0.3299	0.95	0.90	2.95	0.3499
<i>IL22</i>	rs1012356	A/T	12	0.49	3940	All	RD	0.49	683	H	1.08	0.92	1.28	0.3359	1.09	0.93	1.31	0.4368
<i>IL22</i>	rs2227478	A/G	12	0.38	3939	All	RD	0.38	685	R	0.89	0.71	1.13	0.3382	0.90	0.72	1.73	0.24654
<i>IL22</i>	rs2227485	G/A	12	0.42	3928	All	RD	0.43	682	H	1.12	0.95	1.32	0.1632	1.14	0.97	1.33	0.1789
<i>IL22</i>	rs2227491	C/T	12	0.40	3870	All	RD	0.40	672	R	0.92	0.74	1.16	0.4873	0.97	0.81	1.57	0.38947
<i>IL22</i>	rs2227507	T/C	12	0.04	3948	All	RD	0.05	685	H	1.09	0.81	1.46	0.5699	1.10	0.82	1.48	0.4098
<i>TPTE2</i>	rs182873742	C/G	13	0.00	3940	All	RD	0.00	678	H	-	-	-	-	-	-	-	-
<i>RPS6KL1</i>	rs3742785	C/A	14	0.29	3714	All	RD	0.28	656	H	1.06	0.89	1.26	0.4599	1.06	0.89	1.27	0.5001
<i>SPTB</i>	rs229587	T/C	14	0.34	3872	All	RD	0.34	677	R	1.03	0.80	1.32	0.8388	1.09	0.89	1.41	0.10252
<i>YLP1</i>	rs10139016	C/T	14	0.25	3937	All	RD	0.24	684	R	0.88	0.63	1.25	0.4820	0.78	0.37	1.61	0.5046
<i>ADCY9</i>	rs10775349	C/G	16	0.20	3941	All	RD	0.20	601	R	1.26	0.84	1.89	0.2805	1.90	0.97	3.71	0.11955
<i>ADCY9</i>	rs2230739	A/G	16	0.09	3944	All	RD	0.09	602	R	0.64	0.20	2.12	0.4431	0.67	0.32	3.14	0.4588
<i>HBA</i>	α ^{-3.7} -thalassaemia	I/D	16	0.41	3946	All	RD	0.35	662	A	0.78	0.69	0.89	0.0001	0.79	0.70	0.90	0.0002
<i>IL4</i>	rs2243250	T/C	16	0.20	3902	All	RD	0.22	678	H	1.16	0.98	1.38	0.0868	1.21	0.98	1.78	0.1230
<i>ILAR</i>	rs1805015	T/C	16	0.42	3927	All	RD	0.42	683	R	0.87	0.70	1.09	0.2154	1.12	0.81	1.56	0.63147
<i>ADORA2B</i>	rs2535611	T/C	17	0.10	3911	All	RD	0.13	600	A	1.23	1.03	1.49	0.0289	1.28	0.86	1.89	0.2169
<i>NOS2</i>	rs1800482	G/C	17	0.08	3922	All	RD	0.07	679	D	0.94	0.75	1.19	0.6226	0.98	0.56	1.21	0.40375
<i>NOS2</i>	rs2297518	G/A	17	0.14	3910	All	RD	0.14	598	R	1.17	0.65	2.08	0.6096	2.02	0.85	4.79	0.1207
<i>NOS2</i>	rs8078340	C/T	17	0.21	3948	All	RD	0.20	684	R	0.59	0.36	0.99	0.0311	0.61	0.40	0.99	0.032
<i>NOS2</i>	rs9282799	C/T	17	0.03	3949	All	RD	0.03	682	H	1.19	0.84	1.69	0.3324	1.21	0.87	1.72	0.3590
<i>TBX2</i>	rs73991577	A/G	17	0.00	3937	All	RD	0.00	683	H	0.27	0.04	2.05	0.1239	0.30	0.10	2.95	0.21332
<i>EMR1</i>	rs373533	G/T	19	0.46	3863	All	RD	0.47	598	H	1.09	0.92	1.29	0.3338	1.04	0.72	1.51	0.9102
<i>EMR1</i>	rs461645	C/T	19	0.47	3933	All	RD	0.48	601	H	1.11	0.94	1.32	0.2165	1.07	0.73	1.55	0.80482
<i>ICAM</i>	rs5498	A/G	19	0.11	3908	All	RD	0.13	673	H	1.16	0.95	1.41	0.1383	1.16	0.98	3.22	0.3534
<i>GNAS</i>	rs8386	C/T	20	0.26	3914	All	RD	0.29	683	A	1.22	1.07	1.38	0.0028	1.25	1.10	1.43	0.0029
<i>DERL3</i>	rs1128127	G/A	22	0.50	3911	All	RD	0.47	683	A	0.92	0.82	1.03	0.1591	0.88	0.61	1.26	0.3972
<i>CD40LG</i>	rs1126535	T/C	X	0.15	3934	M+F	RD	0.15	683	H	0.82	0.62	1.08	0.1493	0.59	0.37	0.94	0.020226
<i>CD40LG</i>	rs1126535	T/C	X	0.16	1947	F	RD	0.14	326	D	0.82	0.62	1.07	0.1314	0.84	0.63	1.09	0.1414
<i>CD40LG</i>	rs1126535	C/T	X	0.85	1987	M	RD	0.83	357	HM	0.87	0.64	1.19	0.3922	0.88	0.65	1.20	0.41218
<i>CD40LG</i>	rs3092945	T/C	X	0.21	3927	M+F	RD	0.24	680	H	1.28	1.01	1.63	0.0468	1.15	0.81	1.64	0.3059
<i>CD40LG</i>	rs3092945	T/C	X	0.21	1942	F	RD	0.24	325	D	1.29	1.01	1.63	0.0387	1.31	1.03	1.66	0.02
<i>CD40LG</i>	rs3092945	C/T	X	0.78	1985	M	RD	0.77	355	HM	0.95	0.72	1.24	0.7021	0.93	0.71	1.21	0.6100
<i>G6PD</i>	rs1050828	C/T	X	0.19	3940	M+F	RD	0.18	680	D	0.90	0.74	1.09	0.2865	1.05	0.64	1.70	0.64676
<i>G6PD</i>	rs1050828	C/T	X	0.20	1951	F	RD	0.19	325	H	0.89	0.58	1.38	0.6110	0.88	0.68	1.14	0.3500
<i>G6PD</i>	rs1050828	C/T	X	0.19	1989	M	RD	0.17	355	HM	0.89	0.66	1.21	0.8900	0.87	0.64	1.17	0.36
<i>G6PD</i>	rs1050829	T/C	X	0.40	3928	M+F	RD	0.40	681	H	0.88	0.70	1.12	0.2922	0.79	0.51	1.21	0.4246
<i>G6PD</i>	rs1050829	T/C	X	0.41	1952	F	RD	0.40	325	H	0.89	0.70	1.12	0.3148	0.91	0.72	1.16	0.32475
<i>G6PD</i>	rs1050829	C/T	X	0.60	1976	M	RD	0.61	356	HM	1.01	0.80	1.28	0.9095	1.02	0.82	1.30	0.9195

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and rs334 genotype. All individuals having respiratory distress were included in this analysis even if they were identified with multiple severe phenotypes.

Table S6: Odds ratios for all RD for all polymorphisms investigated.

Gene	SNP ID	Alleles (ancestral / derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	MO	Unadjusted				Adjusted [§]			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.08	3940	All	Other SM	0.08	430	R	-	-	-	-	-	-	-	-
AJAP1	rs6674631	G/A	1	0.13	3928	All	Other SM	0.13	431	R	0.89	0.43	1.88	0.7641	0.96	0.46	2.04	0.78675
ATP2B4	rs10900585	G/T	1	0.34	3762	All	Other SM	0.35	426	H	1.19	1.05	1.64	0.0123	1.04	0.63	1.69	0.1783
ATP2B4	rs1541255	A/G	1	0.33	3933	All	Other SM	0.33	431	R	0.84	0.60	1.18	0.3120	0.85	0.60	1.21	0.34498
ATP2B4	rs3753036	G/A	1	0.04	3939	All	Other SM	0.03	432	A	0.82	0.56	1.20	0.2965	0.78	0.52	1.17	0.2440
ATP2B4	rs4951074	G/A	1	0.32	3934	All	Other SM	0.32	431	R	0.89	0.63	1.27	0.5203	1.07	0.73	1.59	0.51529
ATP2B4	rs55868763	G/C	1	0.33	3931	All	Other SM	0.33	431	R	0.85	0.61	1.20	0.3462	0.90	0.62	1.29	0.3814
CR1	rs17047660	A/G	1	0.16	3923	All	Other SM	0.16	436	H	0.85	0.67	1.07	0.1564	0.83	0.65	1.05	0.14157
CR1	rs17047661	G/A	1	0.32	3934	All	Other SM	0.31	434	R	0.91	0.64	1.30	0.6158	0.86	0.59	1.25	0.5502
DARC	rs2814778	G/A	1	0.00	3944	All	Other SM	0.00	379	H	0.58	0.07	4.45	0.5675	0.59	0.08	4.56	0.56766
GBP7	rs1803632	G/C	1	0.49	3942	All	Other SM	0.51	434	R	1.18	0.94	1.47	0.1615	1.12	0.89	1.41	0.1942
IL10	rs1800890	A/T	1	0.24	3945	All	Other SM	0.21	434	R	0.42	0.23	0.76	0.0013	0.43	0.23	0.79	0.001385
IL10	rs1800896	T/C	1	0.38	3943	All	Other SM	0.35	433	R	0.81	0.60	1.11	0.1788	0.91	0.78	1.06	0.1865
IL10	rs3024500	G/A	1	0.45	3928	All	Other SM	0.48	434	R	1.18	0.93	1.50	0.1701	1.18	0.92	1.50	0.17602
LPHN2	rs146428334	A/G	1	0.00	3930	All	Other SM	0.01	430	H	1.55	0.60	4.04	0.3916	1.61	0.61	4.26	0.3839
LPHN2	rs4650365	T/C	1	0.20	3936	All	Other SM	0.17	432	A	0.80	0.66	0.97	0.0183	0.82	0.67	1.00	0.020669
LPHN2	rs72933304	C/A	1	0.09	3942	All	Other SM	0.06	431	D	0.67	0.49	0.91	0.0076	0.68	0.50	0.94	0.0092
LPHN2	rs72933310	T/C	1	0.09	3927	All	Other SM	0.07	431	A	0.71	0.54	0.94	0.0132	0.73	0.55	0.97	0.016043
LPHN2	rs72933350	T/C	1	0.05	3940	All	Other SM	0.03	430	A	0.61	0.41	0.91	0.0095	0.66	0.44	0.99	0.0109
IL1A	rs17561	G/T	2	0.16	3938	All	Other SM	0.19	434	R	1.89	1.16	3.09	0.0162	1.94	1.16	3.22	0.02727
IL1B	rs1143634	C/T	2	0.11	3941	All	Other SM	0.14	434	A	1.20	0.97	1.47	0.0905	1.20	0.95	1.52	0.1054
LAPTM4A	rs973128	C/G	2	0.44	3885	All	Other SM	0.43	419	R	0.86	0.66	1.13	0.2753	0.87	0.66	1.15	0.33759
LOC727982	rs10188961	A/G	2	0.40	3934	All	Other SM	0.44	431	A	1.20	1.04	1.39	0.0123	1.23	1.06	1.43	0.0082
LOC727982	rs1371474	T/C	2	0.27	3933	All	Other SM	0.29	430	D	1.18	0.97	1.45	0.1032	1.22	0.99	1.51	0.077484
LOC727982	rs1371478	C/T	2	0.27	3909	All	Other SM	0.29	429	D	1.18	0.97	1.45	0.0998	1.23	1.00	1.52	0.0751
SDC1	rs11899121	C/G	2	0.47	3921	All	Other SM	0.44	429	R	0.81	0.62	1.04	0.0902	0.84	0.65	1.09	0.11306
ZSWIM2	rs144778284	C/T	2	0.00	3937	All	Other SM	0.00	431	H	0.98	0.29	3.31	0.9741	1.11	0.32	3.80	0.9907
ZSWIM2	rs4316902	A/G	2	0.18	3910	All	Other SM	0.16	430	R	0.42	0.18	0.96	0.0188	0.38	0.16	0.95	0.007496
ARL14	rs74954675	A/C	3	0.08	3907	All	Other SM	0.10	429	A	1.15	0.91	1.47	0.2497	1.17	0.91	1.50	0.2553
ARL14	rs75731597	A/C	3	0.08	3491	All	Other SM	0.10	424	D	1.21	0.93	1.58	0.1547	1.25	0.95	1.64	0.16415
ARL14	rs76033371	A/G	3	0.09	3939	All	Other SM	0.10	431	D	1.14	0.88	1.47	0.3286	1.17	0.90	1.52	0.3341
B3GALNT1	rs12107243	G/C	3	0.49	3934	All	Other SM	0.50	430	R	1.10	0.87	1.38	0.4298	1.16	0.91	1.47	0.34802
IL17RD	rs6780995	A/G	3	0.41	3939	All	Other SM	0.41	433	H	1.27	1.04	1.55	0.0207	1.26	1.03	1.55	0.0164
IL17RE	rs708567	A/G	3	0.50	3881	All	Other SM	0.49	430	R	0.84	0.66	1.07	0.1501	1.14	0.93	1.40	0.14898
TLR9	rs187084	T/C	3	0.33	3933	All	Other SM	0.34	381	H	1.10	0.89	1.36	0.3896	1.13	0.91	1.41	0.3953
FREM3	rs149914432	A/C	4	0.10	3915	All	Other SM	0.06	429	A	0.62	0.47	0.83	0.0006	0.76	0.53	0.86	0.000667
FREM3	rs184895969	C/T	4	0.00	3940	All	Other SM	0.00	431	H	0.50	0.12	2.11	0.2983	0.54	0.12	2.42	0.3061
FREM3	rs184908374	G/A	4	0.00	3943	All	Other SM	0.00	432	H	0.61	0.14	2.59	0.4702	0.67	0.15	3.02	0.48043
FREM3	rs186790584	A/T	4	0.10	3941	All	Other SM	0.06	432	D	0.57	0.42	0.78	0.0002	0.60	0.43	0.8	0.0002
FREM3	rs186873296	A/G	4	0.10	3926	All	Other SM	0.05	428	D	0.55	0.40	0.75	0.0001	0.56	0.42	0.78	0.000195
GAB1	rs7663712	A/G	4	0.02	3938	All	Other SM	0.02	431	R	-	-	-	-	-	-	-	-
INPP4B	rs13103597	C/T	4	0.27	3925	All	Other SM	0.24	431	A	0.88	0.75	1.04	0.1337	0.99	0.82	1.19	0.11272
INPP4B	rs77389579	G/T	4	0.07	3937	All	Other SM	0.04	430	D	0.53	0.37	0.77	0.0003	0.60	0.43	0.8	0.0004
TLR1	rs4833095	C/T	4	0.09	3921	All	Other SM	0.09	380	R	0.61	0.14	2.56	0.4632	0.54	0.13	2.31	0.46587
TLR6	rs5743809	T/C	4	0.06	3884	All	Other SM	0.07	358	A	1.31	0.97	1.76	0.0857	1.35	0.99	1.82	0.0759
TLR6	rs5743810	C/T	4	0.02	3936	All	Other SM	0.02	380	R	-	-	-	-	-	-	-	-
USP38	rs28459062	T/C	4	0.16	3931	All	Other SM	0.17	432	H	1.26	1.01	1.56	0.0417	1.18	0.94	1.48	0.0472
USP38	rs4266246	C/T	4	0.22	3928	All	Other SM	0.21	429	H	0.92	0.75	1.15	0.4720	1.10	0.69	1.74	0.50541
C6	rs1801033	A/C	5	0.47	3927	All	Other SM	0.47	380	H	0.92	0.75	1.14	0.4473	0.93	0.74	1.15	0.3876
IL13	rs20541	C/T	5	0.22	3796	All	Other SM	0.22	360	H	1.01	0.80	1.28	0.9132	1.02	0.81	1.29	0.81604
IRF	rs2706384	C/A	5	0.38	3819	All	Other SM	0.35	365	D	0.89	0.72	1.11	0.3095	0.90	0.72	1.13	0.3295
CTLA4	rs2242665	A/G	6	0.26	3853	All	Other SM	0.32	378	A	1.32	1.12	1.55	0.0011	1.33	1.13	1.58	0.001307
GABBR1	rs192151845	G/A	6	0.01	3943	All	Other SM	0.01	431	H	0.97	0.41	2.27	0.9385	0.86	0.34	2.14	0.9572
HCG4	rs114980857	C/T	6	0.21	3843	All	Other SM	0.22	426	D	1.04	0.85	1.28	0.7093	1.01	0.85	1.21	0.77903
HSPA1B	rs6457452	C/T	6	0.19	3917	All	Other SM	0.16	434	D	0.79	0.64	0.98	0.0332	0.81	0.65	1.01	0.0233
IL20RA	rs1555498	C/T	6	0.46	3947	All	Other SM	0.51	434	D	1.33	1.05	1.68	0.0163	1.29	1.01	1.65	0.016524
LTA	rs2239704	G/T	6	0.12	3896	All	Other SM	0.15	425	H	1.37	1.09	1.72	0.0083	1.28	1.01	1.62	0.0112
LTA	rs909253	T/C	6	0.49	3921	All	Other SM	0.46	428	A	0.92	0.80	1.06	0.2466	0.90	0.70	1.15	0.28612
PLEKHG1	rs114726617	G/A	6	0.04	3940	All	Other SM	0.04	431	R	-	-	-	-	-	-	-	-
PLEKHG1	rs141555199	C/G	6	0.02	3943	All	Other SM	0.03	432	H	1.55	1.02	2.35	0.0514	1.60	1.02	2.49	0.043208
PLEKHG1	rs142712208	G/A	6	0.03	3927	All	Other SM	0.04	431	R	-	-	-	-	-	-	-	-
PLEKHG1	rs15116938	C/T	6	0.03	3939	All	Other SM	0.04	430	R	-	-	-	-	-	-	-	-
PLEKHG1	rs151293197	C/G	6	0.03	3940	All	Other SM	0.03	432	R	-	-	-	-	-	-	-	-
PLEKHG1	rs2131263	C/T	6	0.04	3938	All	Other SM	0.04	430	R	-	-	-	-	-	-	-	-
PLEKHG1	rs55958968	A/G	6	0.05	3932	All	Other SM	0.06	431	H	1.25	0.91	1.72	0.1760	1.22	0.88	1.70	0.1995
PLEKHG1	rs76924464	G/A	6	0.03	3940	All	Other SM	0.03	431	R	-	-	-	-	-	-	-	-
PLEKHG1	rs79100774	G/C	6	0.03	3939	All	Other SM	0.03	432	R	-	-	-	-	-	-	-	-
SNORD48	rs116288147	C/T	6	0.24	3896	All	Other SM	0.21	428	D	0.80	0.65	0.98	0.0331	0.82	0.66	1.01	0.026942
TNF	rs1799964	T/C	6	0.27	3938	All	Other SM	0.27	431	H	0.85	0.69	1.05	0.1238	0.84	0.68	1.04	0.1314
TNF	rs1800629	G/A	6	0.09	3940	All	Other SM	0.08	433	A	0.80	0.61	1.04	0.0862	0.78	0.59	1.03	0.10066
TNF	rs1800750	G/A	6	0.05	3868	All	Other SM	0.05	232	H	0.86	0.53	1.40	0.5408	0.74	0.44	1.24	0.4297
TNF	rs3093662	A/G	6	0.11	3936	All	Other SM	0.11	431	A	0.94	0.75	1.19	0.6219	0.94	0.75	1.19	0.62045
TNF	rs361525	G/A	6	0.07	3948	All	Other SM	0.06	432	H	0.87	0.63	1.19	0.3623	0.82	0.60	1.14	0.3187
CD36	rs3211938	T/G	7	0.10	3942	All	Other SM	0.11	435	A	1.19	0.95	1.49	0.1294	1.24	0.99	1.56	0.13664
NOD1	rs2075820	G/A	7	0.38	3941	All	Other SM											

<i>RRM1</i>	rs116472045	A/G	11	0.02	3939	All	Other SM	0.02	431	H	1.21	0.70	2.10	0.5093	1.22	0.70	2.13	0.47035
<i>RTN3</i>	rs542998	T/C	11	0.43	3929	All	Other SM	0.47	375	D	1.28	1.01	1.62	0.0402	1.31	1.02	1.67	0.0343
<i>TRIM5</i>	rs7935564	G/A	11	0.44	3871	All	Other SM	0.46	378	R	1.21	0.93	1.56	0.1638	1.08	0.83	1.41	0.15605
<i>CAND1</i>	rs10459266	C/A	12	0.21	3938	All	Other SM	0.23	432	D	1.16	0.94	1.42	0.1658	1.17	0.95	1.45	0.1441
<i>CAND1</i>	rs12307123	C/T	12	0.23	3933	All	Other SM	0.22	432	H	0.99	0.80	1.22	0.9170	1.00	0.84	1.19	0.8852
<i>CAND1</i>	rs1566830	G/T	12	0.26	3940	All	Other SM	0.26	431	R	1.14	0.77	1.68	0.5241	1.14	0.76	1.69	0.4937
<i>GRIP1</i>	rs1394263	T/C	12	0.26	3914	All	Other SM	0.26	428	R	1.18	0.80	1.73	0.4130	1.17	0.79	1.73	0.38613
<i>IL22</i>	rs1012356	A/T	12	0.49	3940	All	Other SM	0.48	433	H	1.16	0.95	1.42	0.1362	1.17	0.95	1.44	0.0929
<i>IL22</i>	rs2227478	A/G	12	0.38	3939	All	Other SM	0.40	433	H	0.86	0.70	1.05	0.1377	0.90	0.73	1.11	0.17122
<i>IL22</i>	rs2227485	G/A	12	0.42	3928	All	Other SM	0.41	432	R	0.88	0.67	1.15	0.3423	0.84	0.64	1.12	0.3101
<i>IL22</i>	rs2227491	C/T	12	0.40	3870	All	Other SM	0.43	429	D	1.24	1.00	1.54	0.0520	1.24	0.99	1.56	0.035985
<i>IL22</i>	rs2227507	T/C	12	0.04	3948	All	Other SM	0.04	435	R	0.91	0.11	7.33	0.9290	0.80	0.10	6.56	0.9341
<i>TPTE2</i>	rs182873742	C/G	13	0.00	3940	All	Other SM	0.00	427	H	-	-	-	-	-	-	-	-
<i>RPS6K1</i>	rs3742785	C/A	14	0.29	3714	All	Other SM	0.30	406	H	1.21	0.98	1.49	0.0746	1.14	0.92	1.42	0.1143
<i>SPTB</i>	rs229587	T/C	14	0.34	3872	All	Other SM	0.37	422	R	1.36	1.02	1.81	0.0398	1.34	1.00	1.80	0.043333
<i>YLP1</i>	rs10139016	C/T	14	0.25	3937	All	Other SM	0.26	431	R	1.15	0.78	1.69	0.4848	1.10	0.74	1.64	0.4388
<i>ADCY9</i>	rs10775349	C/G	16	0.20	3941	All	Other SM	0.20	379	H	0.89	0.71	1.12	0.3107	0.87	0.68	1.10	0.30031
<i>ADCY9</i>	rs2230739	A/G	16	0.09	3944	All	Other SM	0.07	380	H	0.72	0.52	1.00	0.0407	0.69	0.50	0.97	0.0431
<i>HBA</i>	α^{37} -thalassaemia	ID	16	0.41	3946	All	Other SM	0.37	421	D	0.80	0.65	0.98	0.0322	0.80	0.64	0.98	0.042911
<i>IL4</i>	rs2243250	T/C	16	0.20	3902	All	Other SM	0.21	428	R	1.12	0.70	1.79	0.6423	1.03	0.86	1.23	0.7517
<i>IL4R</i>	rs1805015	T/C	16	0.42	3927	All	Other SM	0.42	434	A	0.96	0.84	1.11	0.6062	0.99	0.86	1.15	0.71347
<i>ADORA2B</i>	rs2535611	T/C	17	0.10	3911	All	Other SM	0.11	380	R	1.76	0.82	3.80	0.1753	1.76	0.81	3.84	0.1728
<i>NOS2</i>	rs1800482	G/C	17	0.08	3922	All	Other SM	0.08	432	R	0.36	0.05	2.66	0.2383	0.31	0.04	2.40	0.24072
<i>NOS2</i>	rs2297518	G/A	17	0.14	3910	All	Other SM	0.14	377	A	1.02	0.82	1.26	0.8720	1.04	0.84	1.30	0.8338
<i>NOS2</i>	rs8078340	C/T	17	0.21	3948	All	Other SM	0.23	435	R	1.37	0.88	2.13	0.1832	1.28	0.82	2.02	0.17457
<i>NOS2</i>	rs9282799	C/T	17	0.03	3949	All	Other SM	0.02	432	R	-	-	-	-	-	-	-	-
<i>TBX2</i>	rs73991577	A/G	17	0.00	3937	All	Other SM	0.00	430	R	-	-	-	-	-	-	-	-
<i>EMR1</i>	rs373533	G/T	19	0.46	3863	All	Other SM	0.51	379	R	1.31	1.03	1.67	0.0303	1.32	1.03	1.68	0.0260
<i>EMR1</i>	rs461645	C/T	19	0.47	3933	All	Other SM	0.52	380	R	1.33	1.05	1.69	0.0194	1.33	1.04	1.70	0.016471
<i>ICAM</i>	rs5498	A/G	19	0.11	3908	All	Other SM	0.12	427	R	1.12	0.53	2.38	0.7671	1.10	0.51	2.36	0.7497
<i>GNAS</i>	rs8386	C/T	20	0.19	3944	All	Other SM	0.16	434	A	0.84	0.70	1.02	0.0755	0.84	0.69	1.02	0.057606
<i>DERL3</i>	rs1128127	G/A	22	0.50	3911	All	Other SM	0.48	434	D	0.95	0.76	1.18	0.6360	0.98	0.78	1.23	0.6265
<i>CD40LG</i>	rs1126535	T/C	X	0.15	3934	M+F	Other SM	0.15	434	H	0.95	0.68	1.31	0.7387	0.91	0.65	1.28	0.69642
<i>CD40LG</i>	rs1126535	T/C	X	0.16	1947	F	Other SM	0.14	209	R	0.35	0.09	1.47	0.0942	0.37	0.10	1.48	0.0952
<i>CD40LG</i>	rs1126535	C/T	X	0.85	1987	M	Other SM	0.83	225	HM	0.87	0.60	1.26	0.4637	0.89	0.61	1.28	0.4837
<i>CD40LG</i>	rs3092945	T/C	X	0.21	3927	M+F	Other SM	0.26	434	D	1.30	1.05	1.61	0.0165	1.25	1.00	1.56	0.0183
<i>CD40LG</i>	rs3092945	T/C	X	0.21	1942	F	Other SM	0.26	210	D	1.35	1.01	1.80	0.0431	1.36	1.03	1.81	0.0451
<i>CD40LG</i>	rs3092945	C/T	X	0.78	1985	M	Other SM	0.73	224	HM	0.80	0.59	1.10	0.1835	0.81	0.60	1.15	0.2135
<i>G6PD</i>	rs1050828	C/T	X	0.19	3940	M+F	Other SM	0.19	431	H	0.75	0.55	1.04	0.0826	0.73	0.52	1.02	0.082567
<i>G6PD</i>	rs1050828	C/T	X	0.20	1951	F	Other SM	0.17	207	H	0.75	0.55	1.04	0.0827	0.76	0.56	1.06	0.0927
<i>G6PD</i>	rs1050828	C/T	X	0.19	1989	M	Other SM	0.20	224	HM	1.07	0.76	1.52	0.6975	1.09	0.78	1.55	0.7075
<i>G6PD</i>	rs1050829	T/C	X	0.40	3928	M+F	Other SM	0.41	433	H	0.75	0.56	1.01	0.0561	0.83	0.61	1.12	0.0876
<i>G6PD</i>	rs1050829	T/C	X	0.41	1952	F	Other SM	0.41	208	H	0.75	0.56	1.01	0.0555	0.76	0.57	1.02	0.0655
<i>G6PD</i>	rs1050829	C/T	X	0.60	1976	M	Other SM	0.59	225	HM	0.94	0.71	1.24	0.6541	0.95	0.73	1.25	0.6741

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and rs334 genotype. Other SM includes those individuals who did not have CM, SMA or RD.

Table S7: Odds ratios for other SM for all polymorphisms investigated.

Gene	SNPID	Genotype	SM		CM		SMA		RD		Death	
			OR (95% CI)	P [†]	OR (95% CI)	P [†]	OR (95% CI)	P [†]	OR (95% CI)	P [†]	OR (95% CI)	P [†]
<i>ATP2B4</i>	rs1541255	norm (AA)	1		1		1		1		1	
		het (AG)	0.98 (0.88-1.10)	0.8	0.91 (0.80-1.05)	0.22	0.89 (0.75-1.06)	0.22	0.94 (0.79-1.12)	0.53	0.97 (0.74-1.26)	0.83
		hom (GG)	0.73 (0.60-0.89)	0.001	0.65 (0.51-0.83)	0.0007	0.60 (0.44-0.83)	0.002	0.61 (0.44-0.84)	0.002	0.79 (0.49-1.25)	0.32
<i>IL10</i>	rs1800890	norm (AA)	1		1		1		1		1	
		het (AT)	1.03 (0.92-1.15)	0.59	1.01 (0.88-1.15)	0.82	1.06 (0.89-1.26)	0.45	1.03 (0.86-1.22)	0.71	0.90 (0.69-1.19)	0.48
		hom (TT)	0.70 (0.55-0.89)	0.004	0.74 (0.54-1.00)	0.05	0.85 (0.59-1.23)	0.4	0.76 (0.52-1.11)	0.15	0.93 (0.53-1.58)	0.79
<i>LPHN2</i>	rs72933304	norm (CC)	1		1		1		1		1	
		het (AC)	0.82 (0.70-0.95)	0.01	0.84 (0.69-1.01)	0.07	0.86 (0.68-1.09)	0.23	0.84 (0.66-1.06)	0.15	0.79 (0.55-1.15)	0.23
		hom (AA)	0.46 (0.20-1.02)	0.05	0.51 (0.19-1.35)	0.18	0.58 (0.17-1.96)	0.38	0.58 (0.17-1.94)	0.38	-	-
<i>LOC727982</i>	rs1371478	norm (CC)	1		1		1		1		1	
		het (CT)	1.30 (1.16-1.49)	2.70×10 ⁻⁶	1.31 (1.14-1.50)	9.41×10 ⁻⁵	1.34 (1.12-1.59)	0.0008	1.37 (1.15-1.62)	0.0002	1.37 (1.05-1.78)	0.01
		hom (TT)	1.02 (0.83-1.26)	0.78	0.96 (0.74-1.25)	0.79	1.16 (0.85-1.59)	0.35	0.89 (0.63-1.26)	0.53	1.12 (0.69-1.81)	0.62
<i>ARL14</i>	rs75731597	norm (AA)	1		1		1		1		1	
		het (AC)	1.26 (1.09-1.46)	0.001	1.25 (1.04-1.49)	0.01	1.25 (1.02-1.59)	0.03	1.22 (0.98-1.53)	0.06	1.55 (1.13-2.14)	0.006
		hom (CC)	0.80 (0.45-1.43)	0.46	0.73 (0.35-1.55)	0.43	1.07 (0.47-2.44)	0.86	0.73 (0.28-1.88)	0.52	1.23 (0.37-4.07)	0.72
<i>FREM3</i>	rs186873296	norm (AA)	1		1		1		1		1	
		het (AG)	0.57 (0.49-0.68)	3.22×10 ⁻¹¹	0.62 (0.50-0.75)	1.68×10 ⁻⁶	0.56 (0.43-0.73)	2.67×10 ⁻⁵	0.54 (0.41-0.70)	5.76×10 ⁻⁶	0.57 (0.38-0.86)	0.007
		hom (GG)	0.26 (0.11-0.62)	0.002	0.15 (0.03-0.64)	0.01	0.14 (0.01-1.04)	0.05	0.40 (0.12-1.32)	0.13	0.36 (0.04-2.66)	0.31
<i>INPP4B</i>	rs77389579	norm (GG)	1		1		1		1		1	
		het (GT)	0.61 (0.51-0.73)	7.11×10 ⁻⁸	0.64 (0.51-0.80)	0.0001	0.56 (0.41-0.75)	0.0001	0.61 (0.45-0.81)	0.0007	0.51 (0.32-0.83)	0.006
		hom (TT)	0.25 (0.05-1.12)	0.07	0.22 (0.02-1.70)	0.14	0.41 (0.05-3.14)	0.39	-	-	-	-
<i>ABO</i>	rs876719*	non-O	1		1		1		1		1	
		O	0.74 (0.66-0.82)	3.09×10 ⁻⁸	0.78 (0.69-0.89)	0.0003	0.67 (0.57-0.80)	4.13×10 ⁻⁶	0.71 (0.60-0.83)	4.61×10 ⁻⁵	0.67 (0.52-0.87)	0.002
<i>ABO</i>	rs8176746*	norm (CC)	1		1		1		1		1	
		het (AC)	1.35 (1.19-1.52)	9.85×10 ⁻⁷	1.19 (1.03-1.39)	0.01	1.14 (1.17-1.70)	0.0002	1.24 (1.03-1.50)	0.02	1.26 (0.94-1.68)	0.11
		hom (AA)	1.79 (1.29-2.49)	0.0004	1.46 (0.97-2.20)	0.06	2.13 (1.35-3.36)	0.001	2.37 (1.53-3.65)	9.01×10 ⁻⁵	1.61 (0.76-3.41)	0.20
<i>HBB</i>	rs334	norm (AA)	1		1		1		1		1	
		het (AT)	0.15 (0.11-0.20)	2.98×10 ⁻³⁹	0.14 (0.09-0.21)	2.01×10 ⁻²³	0.07 (0.04-0.15)	5.28×10 ⁻¹⁴	0.11 (0.06-0.19)	1.58×10 ⁻¹⁴	0.26 (0.14-0.48)	2.07×10 ⁻⁵
		hom (TT)	0.57 (0.28-1.14)	0.11	0.77 (0.35-1.69)	0.51	1.32 (0.60-2.90)	0.48	1.14 (0.50-2.61)	0.74	1.31 (0.39-4.33)	0.65
<i>RPS6K1</i>	rs3742785	norm (CC)	1		1		1		1		1	
		het (AC)	1.20 (1.07-1.34)	0.001	1.16 (1.01-1.34)	0.03	1.37 (1.15-1.64)	0.0004	1.04 (0.87-1.24)	0.62	0.94 (0.71-1.23)	0.66
		hom (AA)	0.99 (0.81-1.20)	0.93	0.88 (0.69-1.14)	0.35	1.20 (0.89-1.61)	0.2	0.86 (0.63-1.18)	0.37	0.70 (0.42-1.17)	0.18
<i>HBA</i>	α ^{3.7} -thalassaemia	norm	1		1		1		1		1	
		het	0.83 (0.70-0.93)	0.002	0.89 (0.77-1.02)	0.11	0.75 (0.63-0.90)	0.002	0.79 (0.66-0.94)	0.01	0.74 (0.56-0.97)	0.03
		hom	0.66 (0.55-0.78)	1.70×10 ⁻⁶	0.65 (0.52-0.80)	9.68×10 ⁻⁵	0.50 (0.38-0.67)	4.12×10 ⁻⁶	0.61 (0.47-0.81)	0.0004	0.46 (0.29-0.73)	0.0009
<i>CD40LG</i> [§]	rs3092945	norm (TT)	1		1		1		1		1	
		het (CT)	1.12 (0.95-1.13)	0.14	0.98 (0.81-1.20)	0.92	1.18 (0.92-1.51)	0.16	1.28 (1.00-1.64)	0.04	1.30 (0.88-1.92)	0.18
		hom (CC)	1.31 (1.13-1.53)	0.0003	1.32 (1.09-1.59)	0.003	1.37 (1.09-1.72)	0.006	1.07 (0.84-1.37)	0.55	1.00 (0.68-1.46)	0.98
<i>G6PD</i> [§]	rs1050828	norm (CC)	1		1		1		1		1	
		het (CT)	0.82 (0.69-0.96)	0.01	0.79 (0.64-0.97)	0.02	0.80 (0.61-1.04)	0.1	0.88 (0.68-1.13)	0.33	0.60 (0.38-0.94)	0.02
		hom (TT)	1.10 (0.93-1.30)	0.25	0.94 (0.76-1.17)	0.63	1.58 (1.24-2.00)	0.0001	0.92 (0.70-1.21)	0.58	0.69 (0.44-1.09)	0.11

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. †Analyses are adjusted for ethnic group, gender and rs334 genotype. Chr: chromosome; MAF; minor allele frequency; OR: Odds Ratio (95% Confidence Interval); P: p-value. Individuals with multiple phenotypes were included in each of those groups for analysis. §Hemizygous males are coded as homozygotes.

Table S8: Tests of association by specific genotypes for the most significant loci in the case-control study.

Gene	SNPID	Alleles (ancestral/ derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	MO	Unadjusted				Adjusted ^s			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.08	3940	All	CM only	0.08	670	R	1.43	0.58	3.51	0.4556	1.55	0.61	3.94	0.3815
AJAP1	rs6674631	G/A	1	0.13	3928	All	CM only	0.11	666	R	0.49	0.23	1.08	0.0535	0.55	0.25	1.22	0.0571
ATP2B4	rs10900585	G/T	1	0.34	3762	All	CM only	0.34	666	R	1.20	0.90	1.60	0.2112	1.06	0.70	1.59	0.1700
ATP2B4	rs1541255	A/G	1	0.33	3933	All	CM only	0.32	665	R	0.80	0.60	1.06	0.1141	0.81	0.60	1.09	0.0951
ATP2B4	rs3753036	G/A	1	0.04	3939	All	CM only	0.05	668	R	2.46	0.46	13.03	0.3241	1.08	0.80	1.46	0.6336
ATP2B4	rs4951074	G/A	1	0.32	3934	All	CM only	0.31	669	R	0.87	0.65	1.17	0.3565	1.12	0.62	2.03	0.33082
ATP2B4	rs55868763	G/C	1	0.33	3931	All	CM only	0.32	667	R	0.80	0.60	1.07	0.1275	0.82	0.60	1.12	0.1091
CR1	rs17047660	A/G	1	0.16	3923	All	CM only	0.19	659	A	1.18	1.01	1.37	0.0431	1.17	0.99	1.37	0.051
CR1	rs17047661	G/A	1	0.32	3934	All	CM only	0.34	655	A	1.09	0.96	1.24	0.1812	1.18	0.89	1.57	0.1841
DARC	rs2814778	G/A	1	0.00	3944	All	CM only	0.00	583	H	0.71	0.16	3.19	0.6403	0.67	0.15	3.04	0.63892
GBP7	rs1803632	G/C	1	0.49	3942	All	CM only	0.50	662	H	1.03	0.87	1.22	0.7072	1.05	0.86	1.28	0.6713
IL10	rs1800890	A/T	1	0.24	3945	All	CM only	0.22	662	R	0.66	0.44	0.99	0.0372	0.66	0.43	1.00	0.0369
IL10	rs1800896	T/C	1	0.38	3943	All	CM only	0.37	660	H	1.11	0.94	1.31	0.2197	1.10	0.93	1.31	0.2420
IL10	rs3024500	G/A	1	0.45	3928	All	CM only	0.48	661	R	1.16	0.95	1.41	0.1518	1.16	0.94	1.42	0.17422
LPHN2	rs146428334	A/G	1	0.00	3930	All	CM only	0.01	669	H	1.76	0.83	3.75	0.1603	1.87	0.87	4.02	0.1530
LPHN2	rs4650365	T/C	1	0.20	3936	All	CM only	0.19	666	H	0.79	0.66	0.95	0.0107	0.79	0.65	0.95	0.00521
LPHN2	rs72933304	C/A	1	0.09	3942	All	CM only	0.07	669	R	0.19	0.03	1.41	0.0339	0.20	0.03	1.47	0.0350
LPHN2	rs72933310	T/C	1	0.09	3927	All	CM only	0.08	669	A	0.87	0.70	1.07	0.1792	0.90	0.72	1.12	0.18514
LPHN2	rs72933350	T/C	1	0.05	3940	All	CM only	0.04	669	A	0.74	0.55	1.00	0.0447	0.79	0.58	1.08	0.0529
IL1A	rs17561	G/T	2	0.16	3938	All	CM only	0.18	661	R	1.32	0.82	2.12	0.2705	1.41	0.86	2.31	0.2744
IL1B	rs1143634	C/T	2	0.11	3941	All	CM only	0.13	660	R	1.25	0.64	2.44	0.5140	1.40	0.71	2.77	0.5136
LAPTM4A	rs973128	C/G	2	0.44	3885	All	CM only	0.44	661	D	0.99	0.82	1.18	0.8719	0.93	0.78	1.11	0.7362
LOC727982	rs10188961	A/G	2	0.40	3934	All	CM only	0.40	667	D	1.09	0.92	1.30	0.3197	1.13	0.94	1.35	0.3328
LOC727982	rs1371474	T/C	2	0.27	3933	All	CM only	0.29	669	H	1.29	1.09	1.53	0.0031	1.33	1.12	1.58	0.0045
LOC727982	rs1371478	C/T	2	0.27	3909	All	CM only	0.29	668	H	1.30	1.10	1.54	0.0022	1.34	1.13	1.60	0.0033
SDC1	rs11899121	C/G	2	0.47	3921	All	CM only	0.49	666	D	1.15	0.95	1.39	0.1534	1.15	0.95	1.40	0.17548
ZSWIM2	rs144778284	C/T	2	0.00	3937	All	CM only	0.00	669	H	0.21	0.03	1.56	0.0507	0.23	0.03	1.73	0.0518
ZSWIM2	rs4316902	A/G	2	0.18	3910	All	CM only	0.18	667	H	1.04	0.87	1.25	0.6799	1.07	0.89	1.29	0.65552
ARL14	rs74954675	A/C	3	0.08	3907	All	CM only	0.10	662	H	1.32	1.06	1.64	0.0147	1.37	1.10	1.72	0.0109
ARL14	rs75731597	A/C	3	0.08	3491	All	CM only	0.11	657	H	1.34	1.08	1.66	0.0101	1.39	1.11	1.74	0.01012
ARL14	rs76033371	A/G	3	0.09	3939	All	CM only	0.11	665	H	1.25	1.01	1.55	0.0412	1.30	1.04	1.62	0.0377
B3GALNT1	rs12107243	G/C	3	0.49	3934	All	CM only	0.49	668	D	0.94	0.77	1.13	0.4921	0.98	0.86	1.11	0.49661
ILI7RD	rs6780995	A/G	3	0.41	3939	All	CM only	0.39	660	A	0.92	0.82	1.04	0.1948	0.92	0.82	1.04	0.2194
ILI7RE	rs708567	A/G	3	0.50	3881	All	CM only	0.49	660	H	0.98	0.83	1.16	0.8002	0.96	0.81	1.14	0.75464
TLR9	rs187084	T/C	3	0.33	3933	All	CM only	0.31	584	A	0.90	0.79	1.02	0.1075	0.86	0.75	0.99	0.1035
FREM3	rs149914432	A/C	4	0.10	3915	All	CM only	0.05	661	A	0.53	0.41	0.68	7.7x10 ⁻⁰⁸	0.54	0.43	0.70	9.7x10 ⁻⁰⁸
FREM3	rs184895969	C/T	4	0.00	3940	All	CM only	0.00	669	H	0.66	0.23	1.89	0.4161	0.83	0.25	2.70	0.4234
FREM3	rs184908374	G/A	4	0.00	3943	All	CM only	0.00	669	H	0.82	0.28	2.36	0.7000	1.02	0.31	3.39	0.71109
FREM3	rs186790584	A/T	4	0.10	3941	All	CM only	0.05	668	A	0.52	0.41	0.67	6.1x10 ⁻⁰⁸	0.59	0.45	0.8	7.0x10 ⁻⁰⁸
FREM3	rs186873296	A/G	4	0.10	3926	All	CM only	0.05	667	A	0.51	0.39	0.66	2.6x10 ⁻⁰⁸	0.52	0.41	0.67	3.8x10 ⁻⁰⁸
GAB1	rs7663712	A/G	4	0.02	3938	All	CM only	0.03	668	R	-	-	-	-	-	-	-	-
INPP4B	rs13103597	C/T	4	0.27	3925	All	CM only	0.22	668	A	0.76	0.66	0.88	0.0001	0.81	0.69	0.96	0.00016
INPP4B	rs77389579	G/T	4	0.07	3937	All	CM only	0.05	663	A	0.63	0.48	0.83	0.0005	0.69	0.52	0.9	0.0005
TLR1	rs4833095	C/T	4	0.09	3921	All	CM only	0.10	585	R	1.43	0.62	3.29	0.4185	1.31	0.56	3.08	0.41949
TLR6	rs5743809	T/C	4	0.06	3884	All	CM only	0.06	558	R	0.46	0.06	3.51	0.3999	0.45	0.06	3.51	0.3992
TLR6	rs5743810	C/T	4	0.02	3936	All	CM only	0.01	587	A	0.76	0.44	1.32	0.3147	0.61	0.34	1.09	0.31328
USP38	rs28459062	T/C	4	0.16	3931	All	CM only	0.20	669	A	1.26	1.09	1.47	0.0024	1.29	1.08	1.55	0.0029
USP38	rs4266246	C/T	4	0.22	3928	All	CM only	0.22	667	H	1.12	0.94	1.33	0.2077	1.04	0.87	1.25	0.18183
C6	rs1801033	A/C	5	0.47	3927	All	CM only	0.47	584	H	0.97	0.82	1.16	0.7761	1.00	0.83	1.19	0.7991
IL13	rs20541	C/T	5	0.22	3796	All	CM only	0.21	557	R	0.73	0.46	1.14	0.1514	0.70	0.44	1.11	0.17
IRF	rs2706384	C/A	5	0.38	3819	All	CM only	0.36	572	H	0.90	0.75	1.08	0.2449	0.90	0.75	1.08	0.2643
CTLA4	rs2242665	A/G	6	0.26	3853	All	CM only	0.26	575	R	0.91	0.64	1.31	0.6257	0.84	0.58	1.22	0.64253
GABBR1	rs192151845	G/A	6	0.01	3943	All	CM only	0.01	666	H	1.07	0.54	2.12	0.8487	1.12	0.56	2.25	0.8349
HCG4	rs114980857	C/T	6	0.21	3843	All	CM only	0.24	667	D	1.21	1.02	1.43	0.0269	1.19	1.00	1.42	0.026352
HSPA1B	rs6457452	C/T	6	0.19	3917	All	CM only	0.18	658	D	0.89	0.74	1.06	0.1928	0.89	0.74	1.07	0.1974
IL20RA	rs1555498	C/T	6	0.46	3947	All	CM only	0.48	661	H	1.07	0.91	1.27	0.4116	1.09	0.92	1.30	0.40271
LTA	rs2239704	G/T	6	0.12	3896	All	CM only	0.13	653	H	1.09	0.89	1.33	0.3982	1.04	0.85	1.28	0.4347
LTA	rs909253	T/C	6	0.49	3921	All	CM only	0.48	661	H	0.90	0.76	1.06	0.2128	0.92	0.77	1.09	0.21919
PLEKHG1	rs114726617	G/A	6	0.04	3940	All	CM only	0.04	669	A	1.13	0.84	1.51	0.4271	1.17	0.86	1.60	0.4339
PLEKHG1	rs141555199	C/G	6	0.02	3943	All	CM only	0.02	669	R	-	-	-	-	-	-	-	-
PLEKHG1	rs142712208	G/A	6	0.03	3927	All	CM only	0.03	669	D	1.10	0.78	1.54	0.5876	1.19	0.85	1.66	0.6292
PLEKHG1	rs15116938	C/T	6	0.03	3939	All	CM only	0.03	669	A	1.04	0.75	1.44	0.8199	1.13	0.81	1.58	0.86408
PLEKHG1	rs151293197	C/G	6	0.03	3940	All	CM only	0.03	669	R	2.15	0.42	10.94	0.3883	2.83	0.53	15.15	0.3812
PLEKHG1	rs2131263	C/T	6	0.04	3938	All	CM only	0.04	669	A	1.13	0.84	1.51	0.4282	1.17	0.86	1.60	0.435
PLEKHG1	rs55958968	A/G	6	0.05	3932	All	CM only	0.05	669	R	-	-	-	-	-	-	-	-
PLEKHG1	rs76924464	G/A	6	0.03	3940	All	CM only	0.03	669	R	2.15	0.42	10.95	0.3879	2.82	0.53	15.09	0.38075
PLEKHG1	rs79100774	G/C	6	0.03	3939	All	CM only	0.03	670	R	2.15	0.42	10.94	0.3885	2.83	0.53	15.15	0.3813
SNORD48	rs116288147	C/T	6	0.24	3896	All	CM only	0.24	659	H	0.96	0.81	1.14	0.6533	0.97	0.81	1.17	0.75804
TNF	rs1799964	T/C	6	0.27	3938	All	CM only	0.28	654	R	1.11	0.82	1.51	0.5068	1.14	0.83	1.57	0.5155
TNF	rs1800629	G/A	6	0.09	3940	All	CM only	0.08	663	A	0.91	0.73	1.12	0.3730	0.91	0.73	1.13	0.38746
TNF	rs1800750	G/A	6	0.05	3868	All	CM only	0.05	423	R	1.08	0.25	4.70	0.9215	0.97	0.69	1.39	0.9179
TNF	rs3093662	A/G	6	0.11	3936	All	CM only	0.12	659	H	1.10	0.90	1.35	0.3498	1.09	0.89	1.35	0.38042
TNF	rs361525	G/A	6	0.07	3948	All	CM only	0.07	658	R	0.84	0.25	2.83	0.7698				

<i>TRIM5</i>	rs7935564	G/A	11	0.44	3871	All	CM only	0.45	585	H	0.91	0.76	1.08	0.2738	0.90	0.75	1.08	0.29658
<i>CAND1</i>	rs10459266	C/A	12	0.21	3938	All	CM only	0.21	668	R	1.19	0.82	1.73	0.3615	1.18	0.81	1.72	0.3127
<i>CAND1</i>	rs12307123	C/T	12	0.23	3933	All	CM only	0.24	669	A	1.09	0.95	1.25	0.2477	1.07	0.93	1.23	0.25064
<i>CAND1</i>	rs1566830	G/T	12	0.26	3940	All	CM only	0.27	669	A	1.10	0.96	1.25	0.1684	1.08	0.94	1.24	0.1629
<i>GRIP1</i>	rs1394263	T/C	12	0.26	3914	All	CM only	0.27	666	A	1.10	0.97	1.26	0.1508	1.08	0.95	1.24	0.14575
<i>IL22</i>	rs1012356	A/T	12	0.49	3940	All	CM only	0.47	659	D	0.91	0.75	1.09	0.2982	0.93	0.77	1.12	0.2899
<i>IL22</i>	rs2227478	A/G	12	0.38	3939	All	CM only	0.39	661	H	1.02	0.86	1.20	0.8511	1.07	0.90	1.27	0.84254
<i>IL22</i>	rs2227485	G/A	12	0.42	3928	All	CM only	0.40	659	A	0.93	0.83	1.05	0.2441	0.93	0.82	1.05	0.2562
<i>IL22</i>	rs2227491	C/T	12	0.40	3870	All	CM only	0.43	646	A	1.11	0.98	1.25	0.0932	1.09	0.97	1.24	0.090368
<i>IL22</i>	rs2227507	T/C	12	0.04	3948	All	CM only	0.04	663	R	1.79	0.47	6.82	0.4140	1.61	0.41	6.25	0.4152
<i>TPTE2</i>	rs182873742	C/G	13	0.00	3940	All	CM only	0.00	665	H	-	-	-	-	-	-	-	-
<i>RPS6KL1</i>	rs3742785	C/A	14	0.29	3714	All	CM only	0.30	632	H	1.20	1.01	1.43	0.0366	1.17	0.98	1.40	0.0376
<i>SPTB</i>	rs229587	T/C	14	0.34	3872	All	CM only	0.33	642	R	0.78	0.58	1.03	0.0712	0.80	0.60	1.07	0.080607
<i>YLP1</i>	rs10139016	C/T	14	0.25	3937	All	CM only	0.28	668	D	1.27	1.07	1.50	0.0050	1.24	1.04	1.47	0.0049
<i>ADCY9</i>	rs10775349	C/G	16	0.20	3941	All	CM only	0.20	587	R	0.76	0.45	1.27	0.2732	0.81	0.48	1.36	0.27141
<i>ADCY9</i>	rs2230739	A/G	16	0.09	3944	All	CM only	0.07	587	H	0.77	0.60	1.01	0.0492	0.76	0.58	0.99	0.0499
<i>HBA</i>	α ^{3.7} -thalassaemia	I/D	16	0.41	3946	All	CM only	0.37	646	A	0.85	0.75	0.97	0.0126	0.76	0.59	0.99	0.022178
<i>IL4</i>	rs2243250	T/C	16	0.20	3902	All	CM only	0.23	655	D	1.19	1.00	1.41	0.0495	1.17	0.98	1.39	0.0439
<i>IL4R</i>	rs1805015	T/C	16	0.42	3927	All	CM only	0.45	661	D	1.20	1.00	1.44	0.0437	1.24	1.02	1.49	0.040194
<i>ADORA2B</i>	rs2535611	T/C	17	0.10	3911	All	CM only	0.10	586	R	0.28	0.07	1.16	0.0330	0.25	0.06	1.06	0.0325
<i>NOS2</i>	rs1800482	G/C	17	0.08	3922	All	CM only	0.07	659	H	0.90	0.71	1.15	0.4109	0.89	0.69	1.14	0.36808
<i>NOS2</i>	rs2297518	G/A	17	0.14	3910	All	CM only	0.12	582	A	0.86	0.71	1.04	0.1051	0.86	0.71	1.04	0.1030
<i>NOS2</i>	rs8078340	C/T	17	0.21	3948	All	CM only	0.20	662	R	0.70	0.43	1.14	0.1306	0.68	0.42	1.11	0.13073
<i>NOS2</i>	rs9282799	C/T	17	0.03	3949	All	CM only	0.04	659	H	1.44	1.03	2.00	0.0378	1.36	0.95	1.93	0.0484
<i>TBX2</i>	rs73991577	A/G	17	0.00	3937	All	CM only	0.00	669	H	0.79	0.23	2.69	0.6962	0.81	0.23	2.83	0.70518
<i>EMR1</i>	rs373533	G/T	19	0.46	3863	All	CM only	0.50	584	A	1.13	1.00	1.28	0.0503	1.12	0.99	1.28	0.0577
<i>EMR1</i>	rs461645	C/T	19	0.47	3933	All	CM only	0.50	587	A	1.12	0.99	1.27	0.0728	1.11	0.98	1.26	0.082877
<i>ICAM</i>	rs5498	A/G	19	0.11	3908	All	CM only	0.12	652	H	0.99	0.80	1.21	0.9097	0.96	0.78	1.19	0.9401
<i>GNAS</i>	rs8386	C/T	20	0.19	3944	All	CM only	0.20	660	R	1.65	1.11	2.45	0.0186	1.65	1.10	2.48	0.018891
<i>DERL3</i>	rs1128127	G/A	22	0.50	3911	All	CM only	0.48	660	D	0.93	0.77	1.12	0.4317	0.91	0.75	1.11	0.4094
<i>CD40LG</i>	rs1126535	T/C	X	0.15	3934	M+F	CM only	0.17	662	R	1.16	0.87	1.55	0.3039	1.18	0.87	1.58	0.30855
<i>CD40LG</i>	rs1126535	T/C	X	0.16	1947	F	CM only	0.15	330	D	0.92	0.71	1.20	0.5552	0.93	0.73	1.22	0.5652
<i>CD40LG</i>	rs1126535	C/T	X	0.85	1987	M	CM only	0.82	332	HM	0.82	0.60	1.12	0.2124	0.84	0.62	1.22	0.2324
<i>CD40LG</i>	rs3092945	T/C	X	0.21	3927	M+F	CM only	0.26	663	R	1.47	1.17	1.85	0.0013	1.52	1.20	1.92	0.0013
<i>CD40LG</i>	rs3092945	T/C	X	0.21	1942	F	CM only	0.25	331	R	2.27	1.46	3.52	0.0005	2.28	1.48	3.56	0.0005
<i>CD40LG</i>	rs3092945	C/T	X	0.78	1985	M	CM only	0.73	332	HM	0.78	0.60	1.02	0.0736	0.79	0.63	1.09	0.0836
<i>G6PD</i>	rs1050828	C/T	X	0.19	3940	M+F	CM only	0.17	661	D	0.83	0.69	1.02	0.0697	0.85	0.69	1.04	0.069691
<i>G6PD</i>	rs1050828	C/T	X	0.20	1951	F	CM only	0.18	329	H	0.79	0.61	1.02	0.0687	0.80	0.64	1.03	0.0787
<i>G6PD</i>	rs1050828	C/T	X	0.19	1989	M	CM only	0.16	332	HM	0.84	0.61	1.15	0.2796	0.86	0.65	1.16	0.2996
<i>G6PD</i>	rs1050829	T/C	X	0.40	3928	M+F	CM only	0.41	660	H	0.83	0.66	1.05	0.1242	0.87	0.68	1.12	0.1188
<i>G6PD</i>	rs1050829	T/C	X	0.41	1952	F	CM only	0.42	329	H	0.83	0.65	1.05	0.1193	0.85	0.68	1.07	0.1293
<i>G6PD</i>	rs1050829	C/T	X	0.60	1976	M	CM only	0.60	331	HM	0.99	0.78	1.26	0.9445	0.99	0.81	1.28	0.9645

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and rs334 genotype. Only those individuals solely having CM were included in this analysis.

Table S9: Odds ratios for CM only for all polymorphisms investigated.

Gene	SNPID	Alleles (ancestral/derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	M O	Unadjusted				Adjusted [§]			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.92	3940	All	SMA only	0.91	227	R	1.94	0.58	6.45	0.3201	2.00	0.60	6.69	0.3042
AJAP1	rs6674631	G/A	1	0.87	3928	All	SMA only	0.88	228	R	0.91	0.68	1.22	0.5193	0.63	0.20	2.04	0.4130
ATP2B4	rs10900585	G/T	1	0.66	3762	All	SMA only	0.66	226	H	1.16	0.89	1.52	0.2687	1.18	0.90	1.55	0.2251
ATP2B4	rs1541255	A/G	1	0.67	3933	All	SMA only	0.67	226	R	0.86	0.54	1.37	0.5178	0.85	0.53	1.34	0.4698
ATP2B4	rs3753036	G/A	1	0.96	3939	All	SMA only	0.94	228	H	1.46	0.96	2.23	0.0932	1.45	0.95	2.22	0.0993
ATP2B4	rs4951074	G/A	1	0.68	3934	All	SMA only	0.68	227	H	1.17	0.89	1.53	0.2518	1.18	0.90	1.55	0.2202
ATP2B4	rs55868763	G/C	1	0.67	3931	All	SMA only	0.68	225	R	0.83	0.52	1.33	0.4231	0.82	0.51	1.31	0.3877
CR1	rs17047660	A/G	1	0.84	3923	All	SMA only	0.86	230	A	0.85	0.65	1.12	0.2318	0.83	0.63	1.10	0.1810
CR1	rs17047661	G/A	1	0.68	3934	All	SMA only	0.67	229	R	1.16	0.76	1.77	0.4879	1.23	0.81	1.88	0.3447
DARC	rs2814778	G/A	1	1.00	3944	All	SMA only	1.00	207	A	1.47	0.19	11.28	0.7265	1.30	0.17	10.16	0.8073
GBP7	rs1803632	G/C	1	0.51	3942	All	SMA only	0.52	230	H	0.91	0.70	1.19	0.5015	0.91	0.69	1.18	0.4696
IL10	rs1800890	A/T	1	0.76	3945	All	SMA only	0.74	230	H	1.17	0.89	1.53	0.2694	1.20	0.91	1.57	0.2016
IL10	rs1800896	T/C	1	0.62	3943	All	SMA only	0.61	227	R	1.25	0.87	1.78	0.2372	1.28	0.90	1.84	0.1802
IL10	rs3024500	G/A	1	0.55	3928	All	SMA only	0.55	229	H	0.76	0.58	0.99	0.0401	0.75	0.57	0.98	0.0348
LPHN2	rs146428334	A/G	1	1.00	3930	All	SMA only	1.00	228	A	0.52	0.07	3.81	0.4754	0.58	0.08	4.29	0.5627
LPHN2	rs4650365	T/C	1	0.80	3936	All	SMA only	0.81	228	H	0.83	0.62	1.11	0.2109	0.82	0.61	1.10	0.1755
LPHN2	rs72933304	C/A	1	0.91	3942	All	SMA only	0.93	228	R	-	-	-	-	-	-	-	-
LPHN2	rs72933310	T/C	1	0.91	3927	All	SMA only	0.92	228	R	-	-	-	-	-	-	-	-
LPHN2	rs72933350	T/C	1	0.95	3940	All	SMA only	0.96	228	A	0.77	0.47	1.26	0.2768	0.73	0.44	1.20	0.1959
IL1A	rs17561	G/T	2	0.84	3938	All	SMA only	0.85	230	H	0.75	0.55	1.03	0.0697	0.73	0.53	1.01	0.0486
IL1B	rs1143634	C/T	2	0.89	3941	All	SMA only	0.88	230	R	1.77	0.70	4.47	0.2657	1.61	0.63	4.10	0.3464
LAPTM4A	rs973128	C/G	2	0.56	3885	All	SMA only	0.53	225	R	1.28	0.93	1.77	0.1417	1.27	0.92	1.75	0.1620
LOC727982	rs10188961	A/G	2	0.60	3934	All	SMA only	0.58	227	D	1.14	0.86	1.52	0.3559	1.19	0.89	1.58	0.2394
LOC727982	rs1371474	T/C	2	0.73	3933	All	SMA only	0.71	228	H	1.16	0.89	1.53	0.2753	1.19	0.91	1.57	0.2075
LOC727982	rs1371478	C/T	2	0.73	3909	All	SMA only	0.72	226	H	1.14	0.87	1.50	0.3360	1.17	0.89	1.54	0.2564
SDC1	rs11899121	C/G	2	0.53	3921	All	SMA only	0.53	226	H	1.05	0.80	1.37	0.7268	1.06	0.81	1.38	0.6948
ZSWIM2	rs144778284	C/T	2	1.00	3937	All	SMA only	1.00	228	A	1.65	0.38	7.08	0.5285	1.34	0.31	5.78	0.7062
ZSWIM2	rs4316902	A/G	2	0.82	3910	All	SMA only	0.79	226	A	1.20	0.95	1.51	0.1352	1.22	0.96	1.54	0.1087
ARL14	rs74954675	A/C	3	0.92	3907	All	SMA only	0.89	228	H	1.56	1.11	2.18	0.0131	1.46	1.04	2.05	0.0348
ARL14	rs75731597	A/C	3	0.92	3491	All	SMA only	0.90	221	H	1.40	0.99	1.99	0.0649	1.33	0.94	1.89	0.1191
ARL14	rs76033371	A/G	3	0.91	3939	All	SMA only	0.89	228	H	1.48	1.06	2.06	0.0248	1.38	0.99	1.93	0.0636
B3GALNT1	rs12107243	G/C	3	0.51	3934	All	SMA only	0.50	228	D	1.15	0.84	1.58	0.3725	1.13	0.82	1.55	0.4518
IL17RD	rs6780995	A/G	3	0.59	3939	All	SMA only	0.60	230	H	1.23	0.94	1.60	0.1339	1.23	0.94	1.61	0.1288
IL17RE	rs708567	A/G	3	0.50	3881	All	SMA only	0.53	229	R	0.79	0.57	1.09	0.1401	0.80	0.57	1.10	0.1633
TLR9	rs187084	T/C	3	0.67	3933	All	SMA only	0.67	208	R	0.94	0.60	1.46	0.7783	0.92	0.59	1.44	0.7131
FREM3	rs149914432	A/C	4	0.90	3915	All	SMA only	0.94	226	A	0.56	0.38	0.84	0.0024	0.56	0.38	0.84	0.0026
FREM3	rs184895969	C/T	4	1.00	3940	All	SMA only	1.00	228	A	1.12	0.27	4.69	0.8829	1.01	0.24	4.25	0.9932
FREM3	rs184908374	G/A	4	1.00	3943	All	SMA only	1.00	228	A	1.39	0.33	5.89	0.6719	1.23	0.29	5.27	0.7822
FREM3	rs186790584	A/T	4	0.90	3941	All	SMA only	0.95	228	A	0.54	0.36	0.82	0.0015	0.55	0.36	0.83	0.0018
FREM3	rs186873296	A/G	4	0.90	3926	All	SMA only	0.95	226	A	0.51	0.33	0.78	0.0006	0.51	0.33	0.78	0.0007
GAB1	rs7663712	A/G	4	0.98	3938	All	SMA only	0.98	228	A	0.79	0.40	1.55	0.4748	0.82	0.42	1.62	0.5562
INPP4B	rs13103597	C/T	4	0.73	3925	All	SMA only	0.77	228	D	0.83	0.66	1.03	0.0912	0.79	0.60	1.04	0.0849
INPP4B	rs77389579	G/T	4	0.93	3937	All	SMA only	0.95	228	A	0.65	0.41	1.01	0.0430	0.64	0.41	1.00	0.0383
TLR1	rs4833095	C/T	4	0.91	3921	All	SMA only	0.92	208	H	0.65	0.42	1.05	0.0643	0.69	0.45	1.06	0.0743
TLR6	rs5743809	T/C	4	0.94	3884	All	SMA only	0.96	200	H	0.65	0.37	1.13	0.1036	0.63	0.36	1.10	0.0822
TLR6	rs5743810	C/T	4	0.98	3936	All	SMA only	0.98	208	R	-	-	-	-	-	-	-	-
USP38	rs28459062	T/C	4	0.84	3931	All	SMA only	0.83	227	H	1.06	0.79	1.43	0.6870	1.06	0.79	1.44	0.6864
USP38	rs4266246	C/T	4	0.78	3928	All	SMA only	0.76	227	R	1.85	1.13	3.02	0.0227	1.89	1.15	3.09	0.0195
C6	rs1801033	A/C	5	0.53	3927	All	SMA only	0.53	207	H	0.94	0.71	1.24	0.6586	0.92	0.70	1.22	0.5694
IL13	rs20541	C/T	5	0.78	3796	All	SMA only	0.78	195	R	0.48	0.20	1.19	0.0763	0.46	0.19	1.13	0.0560
IRF	rs2706384	C/A	5	0.62	3819	All	SMA only	0.65	204	R	0.67	0.42	1.06	0.0695	0.68	0.43	1.08	0.0904
CTL4	rs2242665	A/G	6	0.74	3853	All	SMA only	0.78	204	D	0.69	0.52	0.92	0.0115	0.69	0.52	0.93	0.0131
GABBR1	rs192151845	G/A	6	0.99	3943	All	SMA only	1.00	227	A	0.31	0.04	2.23	0.1529	0.31	0.04	2.27	0.1612
HCG4	rs114980857	C/T	6	0.79	3843	All	SMA only	0.79	225	R	0.70	0.34	1.45	0.3171	0.73	0.35	1.49	0.3625
HSPA1B	rs6457452	C/T	6	0.81	3917	All	SMA only	0.84	230	A	0.79	0.61	1.02	0.0675	0.79	0.61	1.02	0.0586
IL20RA	rs1555498	C/T	6	0.54	3947	All	SMA only	0.48	229	R	1.22	1.01	1.47	0.0363	1.31	0.97	1.77	0.0824
LTA	rs2239704	G/T	6	0.88	3896	All	SMA only	0.90	223	R	-	-	-	-	-	-	-	-
LTA	rs909253	T/C	6	0.51	3921	All	SMA only	0.46	227	D	1.38	0.99	1.92	0.0502	1.42	1.02	1.98	0.0340
PLEKHG1	rs114726617	G/A	6	0.96	3940	All	SMA only	0.96	228	R	2.17	0.27	17.39	0.5100	2.09	0.26	16.98	0.5279
PLEKHG1	rs141555199	C/G	6	0.98	3943	All	SMA only	0.98	228	A	0.76	0.35	1.62	0.4582	0.71	0.33	1.52	0.3470
PLEKHG1	rs142712208	G/A	6	0.97	3927	All	SMA only	0.97	228	R	2.88	0.35	24.02	0.3888	3.10	0.37	26.17	0.3621
PLEKHG1	rs15116938	C/T	6	0.97	3939	All	SMA only	0.96	228	R	2.89	0.35	24.09	0.3876	3.11	0.37	26.23	0.3611
PLEKHG1	rs151293197	C/G	6	0.97	3940	All	SMA only	0.97	228	R	2.89	0.35	24.09	0.3875	3.11	0.37	26.24	0.3610
PLEKHG1	rs2131263	C/T	6	0.96	3938	All	SMA only	0.96	228	R	2.16	0.27	17.38	0.5103	2.09	0.26	16.97	0.5281
PLEKHG1	rs55958968	A/G	6	0.95	3932	All	SMA only	0.94	228	A	1.23	0.83	1.81	0.3141	1.28	0.87	1.90	0.2221
PLEKHG1	rs76924464	G/A	6	0.97	3940	All	SMA only	0.97	228	R	2.89	0.35	24.09	0.3875	3.11	0.37	26.25	0.3609
PLEKHG1	rs79100774	G/C	6	0.97	3939	All	SMA only	0.97	228	R	2.89	0.35	24.09	0.3876	3.11	0.37	26.23	0.3612
SNORD48	rs116288147	C/T	6	0.76	3896	All	SMA only	0.79	227	A	0.85	0.67	1.07	0.1539	0.82	0.65	1.03	0.0855
TNF	rs1799964	T/C	6	0.73	3938	All	SMA only	0.71	228	R	1.05	0.86	1.30	0.6195	1.13	0.70	1.84	0.6212
TNF	rs1800629	G/A	6	0.91	3940	All	SMA only	0.90	228	R	1.74	0.53	5.74	0.3991	1.81	0.54	5.99	0.3711
TNF	rs1800750	G/A	6	0.95	3868	All	SMA only	0.92	125	H	1.62	0.98	2.66	0.0734	1.71	1.04	2.83	0.0474
TNF	rs3093662	A/G	6	0.89	3936	All	SMA only	0.90	227	H	0.79	0.55	1.14	0.1984	0.77	0.53	1.11	0.1531
TNF	rs361525	G/A	6	0.93	3948	All	SMA only	0.93	229	R	-	-	-	-	-	-	-	-
CD36	rs3211938	T/G	7	0.90	3942	All	SMA only	0.91	230	R	0.42							

MKI67	rs115947774	A/G	10	1.00	3931	All	SMA only	1.00	226	A	0.67	0.09	4.94	0.6734	0.57	0.08	4.21	0.5448
MKI67	rs148494166	T/C	10	1.00	3913	All	SMA only	1.00	214	A	-	-	-	-	-	-	-	-
HBB	rs334	A/T	11	0.92	3949	All	SMA only	0.98	228	H	0.10	0.04	0.27	9.68×10 ⁻¹²	0.10	0.04	0.28	3.87×10 ⁻¹¹
RRM1	rs116472045	A/G	11	0.98	3939	All	SMA only	0.97	228	H	1.98	1.10	3.56	0.0368	2.01	1.11	3.64	0.0334
RTN3	rs542998	T/C	11	0.57	3929	All	SMA only	0.51	201	D	1.28	1.05	1.56	0.0164	1.44	1.04	2.00	0.0234
TRIM5	rs7935564	G/A	11	0.56	3871	All	SMA only	0.51	206	D	1.47	1.06	2.05	0.0177	1.44	1.03	2.00	0.0269
CAND1	rs10459266	C/A	12	0.79	3938	All	SMA only	0.76	228	H	1.31	1.00	1.73	0.0547	1.32	1.00	1.74	0.0522
CAND1	rs12307123	C/T	12	0.77	3933	All	SMA only	0.78	228	H	0.83	0.62	1.11	0.2087	0.84	0.63	1.12	0.2402
CAND1	rs1566830	G/T	12	0.74	3940	All	SMA only	0.74	227	H	0.84	0.64	1.11	0.2251	0.84	0.64	1.12	0.2338
GRIP1	rs1394263	T/C	12	0.74	3914	All	SMA only	0.74	225	H	0.83	0.62	1.10	0.1805	0.83	0.62	1.10	0.1887
IL22	rs1012356	A/T	12	0.51	3940	All	SMA only	0.50	229	D	1.07	0.82	1.40	0.6041	1.10	0.80	1.50	0.5615
IL22	rs2227478	A/G	12	0.62	3939	All	SMA only	0.60	230	D	1.19	0.90	1.57	0.2291	1.14	0.86	1.51	0.3608
IL22	rs2227485	G/A	12	0.58	3928	All	SMA only	0.60	226	R	0.67	0.45	1.00	0.0380	0.69	0.46	1.03	0.0602
IL22	rs2227491	C/T	12	0.60	3870	All	SMA only	0.62	227	R	0.69	0.46	1.05	0.0695	0.68	0.45	1.03	0.0549
IL22	rs2227507	T/C	12	0.96	3948	All	SMA only	0.96	230	R	-	-	-	-	-	-	-	-
TPTE2	rs182873742	C/G	13	1.00	3940	All	SMA only	1.00	228	A	-	-	-	-	-	-	-	-
RPS6KL1	rs3742785	C/A	14	0.71	3714	All	SMA only	0.66	222	D	1.38	1.05	1.81	0.0212	1.40	1.06	1.84	0.0156
SPTB	rs229587	T/C	14	0.66	3872	All	SMA only	0.66	210	H	0.90	0.68	1.20	0.4737	0.89	0.67	1.18	0.4127
YLPM1	rs10139016	C/T	14	0.75	3937	All	SMA only	0.70	227	D	1.33	1.02	1.74	0.0381	1.36	1.04	1.78	0.0243
ADCY9	rs10775349	C/G	16	0.80	3941	All	SMA only	0.79	207	H	1.08	0.81	1.45	0.6033	1.08	0.81	1.46	0.5945
ADCY9	rs2230739	A/G	16	0.91	3944	All	SMA only	0.91	208	H	0.61	0.08	4.48	0.5993	1.08	0.75	1.58	0.6750
HBA	α ^{-3.7} -thalassaemia	I/D	16	0.59	3946	All	SMA only	0.67	218	A	0.71	0.57	0.87	0.0009	0.70	0.57	0.86	0.0007
IL4	rs2243250	T/C	16	0.80	3902	All	SMA only	0.77	224	H	1.31	1.00	1.72	0.0560	1.30	0.98	1.71	0.0716
IL4R	rs1805015	T/C	16	0.58	3927	All	SMA only	0.59	229	D	0.91	0.69	1.20	0.5018	0.90	0.68	1.19	0.4620
ADORA2B	rs2535611	T/C	17	0.90	3911	All	SMA only	0.90	207	R	0.45	0.06	3.27	0.3649	0.40	0.05	2.94	0.2958
NOS2	rs1800482	G/C	17	0.92	3922	All	SMA only	0.91	229	H	1.18	0.83	1.69	0.3695	1.20	0.84	1.72	0.3328
NOS2	rs2297518	G/A	17	0.86	3910	All	SMA only	0.85	205	A	1.09	0.83	1.44	0.5371	1.07	0.81	1.42	0.6169
NOS2	rs8078340	C/T	17	0.79	3948	All	SMA only	0.80	230	H	0.82	0.61	1.10	0.1803	0.83	0.62	1.11	0.2009
NOS2	rs9282799	C/T	17	0.97	3949	All	SMA only	0.97	229	H	1.34	0.78	2.30	0.3142	1.29	0.75	2.22	0.3804
TBX2	rs73991577	A/G	17	1.00	3937	All	SMA only	1.00	228	A	0.91	0.12	6.82	0.9245	0.78	0.10	5.91	0.8061
EMR1	rs373533	G/T	19	0.54	3863	All	SMA only	0.51	207	R	1.19	0.86	1.64	0.3115	1.15	0.83	1.59	0.4097
EMR1	rs461645	C/T	19	0.53	3933	All	SMA only	0.51	208	R	1.19	0.86	1.64	0.3071	1.15	0.83	1.59	0.4081
ICAM	rs5498	A/G	19	0.89	3908	All	SMA only	0.88	226	R	1.91	0.81	4.49	0.1707	1.73	0.73	4.09	0.2410
GNAS	rs8386	C/T	20	0.81	3944	All	SMA only	0.79	230	R	1.68	0.92	3.09	0.1159	1.69	0.92	3.10	0.1153
DERL3	rs1128127	G/A	22	0.50	3911	All	SMA only	0.57	228	A	0.78	0.64	0.94	0.0076	0.79	0.65	0.96	0.0154
CD40LG	rs1126535	T/C	x	0.85	3934	M+F	SMA only	0.86	229	H	0.60	0.37	0.98	0.0280	0.68	0.41	1.14	0.1355
CD40LG	rs1126535	T/C	x	0.85	3934	F	SMA only	0.86	229	H	0.70	0.42	1.17	0.1605	0.73	0.47	1.14	0.1341
CD40LG	rs1126535	T/C	x	0.85	3934	M	SMA only	0.86	229	HM	0.93	0.57	1.50	0.7596	0.94	0.58	1.52	0.7940
CD40LG	rs3092945	T/C	x	0.79	3927	M+F	SMA only	0.69	227	A	1.94	1.41	2.68	0.0001	1.35	1.14	1.59	0.0005
CD40LG	rs3092945	T/C	x	0.79	3927	F	SMA only	0.69	227	H	1.78	1.30	2.43	0.0005	1.74	1.27	2.39	0.0008
CD40LG	rs3092945	T/C	x	0.79	3927	M	SMA only	0.69	227	HM	0.65	0.44	0.95	0.0321	0.66	0.45	0.97	0.0411
G6PD	rs1050828	C/T	x	0.81	3940	M+F	SMA only	0.76	229	R	1.74	1.23	2.48	0.0033	1.63	1.13	2.36	0.0122
G6PD	rs1050828	C/T	x	0.81	3940	F	SMA only	0.76	229	H	0.85	0.54	1.33	0.4602	0.84	0.54	1.32	0.4465
G6PD	rs1050828	C/T	x	0.81	3940	M	SMA only	0.76	229	HM	1.72	1.16	2.54	0.0093	1.76	1.18	2.60	0.0070
G6PD	rs1050829	T/C	x	0.60	3928	M+F	SMA only	0.57	229	R	0.80	0.57	1.11	0.1659	1.12	0.83	1.51	0.4527
G6PD	rs1050829	T/C	x	0.60	3928	F	SMA only	0.57	229	H	0.84	0.62	1.13	0.2401	0.83	0.62	1.13	0.2426
G6PD	rs1050829	T/C	x	0.60	3928	M	SMA only	0.57	229	HM	0.75	0.53	1.07	0.1164	0.76	0.53	1.08	0.1282

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and *rs334*. Only those individuals solely having SMA were included in this analysis.

Table S10: Odds ratios for SMA only for all polymorphisms investigated.

Gene	SNPID	Alleles (ancestral/ derived)	Chr	DAF Controls	No. of genotype in controls	Sample	Case Phenotype	DAF Cases	No. of genotypes in cases	MO	Unadjusted				Adjusted ^s			
											OR	LCL	UCL	P	OR	LCL	UCL	P
AJAP1	rs113788643	A/G	1	0.08	3940	All	RD_only	0.06	157	R	-	-	-	-	-	-	-	-
AJAP1	rs6674631	G/A	1	0.13	3928	All	RD_only	0.14	157	H	1.28	0.89	1.85	0.1872	1.33	0.92	1.93	0.15938
ATP2B4	rs10900585	G/T	1	0.34	3762	All	RD_only	0.37	157	H	1.66	1.05	2.85	0.0229	1.67	1.08	2.44	0.0251
ATP2B4	rs1541255	A/G	1	0.33	3933	All	RD_only	0.36	157	H	0.63	0.48	0.87	0.0275	0.66	0.50	0.90	0.027579
ATP2B4	rs3753036	G/A	1	0.04	3939	All	RD_only	0.04	157	R	4.94	0.57	42.62	0.2241	7.94	0.83	76.21	0.2218
ATP2B4	rs4951074	G/A	1	0.32	3934	All	RD_only	0.35	157	H	1.32	0.96	1.82	0.0879	1.09	0.59	2.04	0.063237
ATP2B4	rs55868763	G/C	1	0.33	3931	All	RD_only	0.36	157	H	1.39	1.01	1.92	0.0432	1.49	1.10	1.98	0.0440
CR1	rs17047660	A/G	1	0.16	3923	All	RD_only	0.18	156	A	1.15	0.85	1.55	0.3637	1.22	0.90	1.66	0.33548
CR1	rs17047661	G/A	1	0.32	3934	All	RD_only	0.33	155	R	1.37	0.85	2.22	0.2151	1.22	0.73	2.03	0.2094
DARC	rs2814778	G/A	1	0.00	3944	All	RD_only	0.00	151	H	-	-	-	-	-	-	-	-
GBP7	rs1803632	G/C	1	0.49	3942	All	RD_only	0.49	156	H	0.85	0.61	1.17	0.3037	0.89	0.64	1.23	0.3454
IL10	rs1800890	A/T	1	0.24	3945	All	RD_only	0.22	156	R	0.29	0.09	0.91	0.0092	0.27	0.08	0.88	0.009432
IL10	rs1800896	T/C	1	0.38	3943	All	RD_only	0.36	156	R	0.75	0.45	1.25	0.2542	0.74	0.44	1.25	0.2693
IL10	rs3024500	G/A	1	0.45	3928	All	RD_only	0.45	155	H	1.35	0.97	1.87	0.0707	1.36	0.98	1.90	0.057556
LPHN2	rs146428334	A/G	1	0.00	3930	All	RD_only	0.01	157	H	1.53	0.36	6.43	0.5876	1.54	0.36	6.61	0.5752
LPHN2	rs4650365	T/C	1	0.20	3936	All	RD_only	0.19	157	R	0.70	0.25	1.91	0.4562	0.77	0.28	2.11	0.48287
LPHN2	rs72933304	C/A	1	0.09	3942	All	RD_only	0.09	157	H	1.12	0.73	1.70	0.6173	1.14	0.74	1.76	0.5766
LPHN2	rs72933310	T/C	1	0.09	3927	All	RD_only	0.10	157	H	1.13	0.75	1.71	0.5678	1.14	0.75	1.74	0.52604
LPHN2	rs72933350	T/C	1	0.05	3940	All	RD_only	0.07	157	A	1.40	0.89	2.22	0.1663	1.47	0.92	2.35	0.1538
IL1A	rs17561	G/T	2	0.16	3938	All	RD_only	0.20	156	R	2.15	1.02	4.51	0.0664	2.39	1.13	5.09	0.064612
IL1B	rs1143634	C/T	2	0.11	3941	All	RD_only	0.10	156	A	0.84	0.57	1.22	0.3494	0.81	0.55	1.20	0.3628
LAPTM4A	rs973128	C/G	2	0.44	3885	All	RD_only	0.35	156	D	0.55	0.40	0.75	0.0003	0.53	0.38	0.74	0.000191
LOC727982	rs10188961	A/G	2	0.40	3934	All	RD_only	0.44	157	A	1.20	0.95	1.50	0.1242	1.17	0.93	1.47	0.1616
LOC727982	rs1371474	T/C	2	0.27	3933	All	RD_only	0.31	157	H	1.63	1.18	2.25	0.0029	1.69	1.22	2.35	0.002967
LOC727982	rs1371478	C/T	2	0.27	3909	All	RD_only	0.30	157	H	1.71	1.24	2.36	0.0011	1.73	1.25	2.40	0.0019
SDC1	rs11899121	C/G	2	0.47	3921	All	RD_only	0.50	157	D	1.24	0.85	1.80	0.2514	1.27	0.87	1.85	0.29507
ZSWIM2	rs144778284	C/T	2	0.00	3937	All	RD_only	0.00	157	H	1.13	0.15	8.51	0.9058	1.31	0.17	10.14	0.9004
ZSWIM2	rs4316902	A/G	2	0.18	3910	All	RD_only	0.17	153	H	0.75	0.51	1.10	0.1343	0.79	0.53	1.16	0.16013
ARL14	rs74954675	A/C	3	0.08	3907	All	RD_only	0.08	156	R	-	-	-	-	-	-	-	-
ARL14	rs75731597	A/C	3	0.08	3491	All	RD_only	0.10	157	R	-	-	-	-	-	-	-	-
ARL14	rs76033371	A/G	3	0.09	3939	All	RD_only	0.10	157	R	-	-	-	-	-	-	-	-
B3GALNT1	rs12107243	G/C	3	0.49	3934	All	RD_only	0.46	157	A	0.89	0.71	1.13	0.3422	0.93	0.74	1.18	0.39756
IL17RD	rs6780995	A/G	3	0.41	3939	All	RD_only	0.40	156	R	0.96	0.62	1.47	0.8356	0.98	0.63	1.51	0.8753
IL17RE	rs708567	A/G	3	0.50	3881	All	RD_only	0.45	154	A	0.82	0.66	1.04	0.0967	0.81	0.64	1.02	0.11682
TLR9	rs187084	T/C	3	0.33	3933	All	RD_only	0.33	151	R	0.83	0.48	1.43	0.4990	1.14	0.81	1.58	0.4593
FREM3	rs149914432	A/C	4	0.10	3915	All	RD_only	0.06	157	H	0.46	0.26	0.80	0.0025	0.48	0.24	0.95	0.002561
FREM3	rs184895969	C/T	4	0.00	3940	All	RD_only	0.01	157	H	3.13	1.09	9.02	0.0649	8.15	2.19	30.36	0.0633
FREM3	rs184908374	G/A	4	0.00	3943	All	RD_only	0.01	157	H	2.90	0.86	9.74	0.1313	6.51	1.54	27.44	0.12848
FREM3	rs186790584	A/T	4	0.10	3941	All	RD_only	0.06	157	H	0.47	0.27	0.81	0.0031	0.47	0.27	0.8	0.0033
FREM3	rs186873296	A/G	4	0.10	3926	All	RD_only	0.06	157	H	0.47	0.27	0.83	0.0038	0.48	0.29	0.82	0.003987
GAB1	rs7663712	A/G	4	0.02	3938	All	RD_only	0.01	157	A	0.51	0.19	1.39	0.1433	0.51	0.19	1.38	0.1530
INPP4B	rs13103597	C/T	4	0.27	3925	All	RD_only	0.21	157	D	0.64	0.46	0.90	0.0086	0.70	0.48	1.02	0.004952
INPP4B	rs77389579	G/T	4	0.07	3937	All	RD_only	0.04	157	A	0.57	0.32	1.01	0.0358	0.70	0.46	0.9	0.0213
TLR1	rs4833095	C/T	4	0.09	3921	All	RD_only	0.12	150	A	1.40	0.98	2.01	0.0764	1.31	0.90	1.89	0.070727
TLR6	rs5743809	T/C	4	0.06	3884	All	RD_only	0.06	145	R	3.68	0.82	16.40	0.1441	3.71	0.81	17.04	0.1424
TLR6	rs5743810	C/T	4	0.02	3936	All	RD_only	0.03	151	H	2.02	1.00	4.09	0.0737	2.02	0.99	4.14	0.071669
USP38	rs28459062	T/C	4	0.16	3931	All	RD_only	0.18	157	D	1.11	0.79	1.57	0.5522	1.05	0.78	1.43	0.5949
USP38	rs4266246	C/T	4	0.22	3928	All	RD_only	0.22	157	R	1.42	0.73	2.74	0.3195	1.34	0.69	2.61	0.29847
C6	rs1801033	A/C	5	0.47	3927	All	RD_only	0.44	150	D	0.72	0.51	1.02	0.0654	0.72	0.51	1.02	0.0586
IL13	rs20541	C/T	5	0.22	3796	All	RD_only	0.24	144	A	1.13	0.86	1.48	0.3936	1.13	0.86	1.48	0.34347
IRF	rs2706384	C/A	5	0.38	3819	All	RD_only	0.38	144	R	1.13	0.72	1.79	0.6008	1.06	0.67	1.69	0.5697
CTLA	rs2242665	A/G	6	0.26	3853	All	RD_only	0.32	150	R	1.83	1.10	3.05	0.0289	1.32	1.02	1.70	0.025206
GABBR1	rs192151845	G/A	6	0.01	3943	All	RD_only	0.01	157	H	1.33	0.41	4.32	0.6438	0.88	0.21	3.67	0.8750
HCG4	rs114980857	C/T	6	0.21	3843	All	RD_only	0.20	150	A	0.91	0.69	1.22	0.5392	0.89	0.67	1.19	0.57831
HSPA1B	rs6457452	C/T	6	0.19	3917	All	RD_only	0.17	157	A	0.86	0.63	1.16	0.3047	0.85	0.63	1.16	0.2810
IL20RA	rs1555498	C/T	6	0.46	3947	All	RD_only	0.44	156	D	0.82	0.58	1.15	0.2508	0.87	0.61	1.24	0.30236
LTA	rs2239704	G/T	6	0.12	3896	All	RD_only	0.14	153	H	1.37	0.95	1.97	0.0955	1.32	0.91	1.91	0.0873
LTA	rs909253	T/C	6	0.49	3921	All	RD_only	0.48	155	D	0.96	0.67	1.38	0.8402	1.00	0.79	1.25	0.91093
PLEKHG1	rs114726617	G/A	6	0.04	3940	All	RD_only	0.05	157	H	1.41	0.81	2.43	0.2391	1.37	0.77	2.44	0.3355
PLEKHG1	rs141555199	C/G	6	0.02	3943	All	RD_only	0.02	157	R	-	-	-	-	-	-	-	-
PLEKHG1	rs142712208	G/A	6	0.03	3927	All	RD_only	0.04	157	H	1.29	0.70	2.36	0.4273	1.38	0.74	2.57	0.3974
PLEKHG1	rs15116938	C/T	6	0.03	3939	All	RD_only	0.04	157	R	-	-	-	-	-	-	-	-
PLEKHG1	rs151293197	C/G	6	0.03	3940	All	RD_only	0.04	156	H	1.25	0.68	2.29	0.4826	1.32	0.71	2.46	0.4501
PLEKHG1	rs2131263	C/T	6	0.04	3938	All	RD_only	0.05	157	H	1.41	0.81	2.43	0.2397	1.37	0.77	2.44	0.33629
PLEKHG1	rs55958968	A/G	6	0.05	3932	All	RD_only	0.07	157	H	1.42	0.89	2.29	0.1578	1.39	0.85	2.25	0.1400
PLEKHG1	rs76924464	G/A	6	0.03	3940	All	RD_only	0.04	157	R	-	-	-	-	-	-	-	-
PLEKHG1	rs79100774	G/C	6	0.03	3939	All	RD_only	0.04	157	R	-	-	-	-	-	-	-	-
SNORD48	rs116288147	C/T	6	0.24	3896	All	RD_only	0.21	156	A	0.86	0.65	1.14	0.2896	0.55	0.22	1.39	0.17612
TNF	rs1799964	T/C	6	0.27	3938	All	RD_only	0.24	155	R	0.58	0.27	1.25	0.1315	0.50	0.22	1.14	0.0640
TNF	rs1800629	G/A	6	0.09	3940	All	RD_only	0.09	156	R	3.46	1.20	9.97	0.0477	3.37	1.13	9.98	0.04709
TNF	rs1800750	G/A	6	0.05	3868	All	RD_only	0.03	58	A	0.66	0.24	1.77	0.3711	0.67	0.25	1.81	0.3678
TNF	rs3093662	A/G	6	0.11	3936	All	RD_only	0.10	154	D	0.87	0.58	1.32	0.5198	0.87	0.57	1.32	0.54979
TNF	rs361525	G/A	6	0.07	3948	All	RD_only	0.05	156	R	-	-	-	-	-	-	-	-
CD36	rs3211938	T/G	7	0.10	3942	All	RD_only	0.09	155	H	0.88	0.56	1.37	0.5574	0.89	0.57	1.40	0.57785
NOD1	rs2075820	G/A	7	0.38	3941	All	RD											

<i>MKI67</i>	rs11016116	A/G	10	0.02	3944	All	RD_only	0.03	157	R	-	-	-	-	-	-	-	-
<i>MKI67</i>	rs115947774	A/G	10	0.00	3931	All	RD_only	0.00	156	H	0.91	0.12	6.79	0.9269	0.79	0.10	5.98	0.9345
<i>MKI67</i>	rs148494166	T/C	10	0.00	3913	All	RD_only	0.01	146	H	3.58	0.81	15.78	0.1496	3.30	0.74	14.82	0.14532
<i>HBB</i>	rs334	A/T	11	0.08	3949	All	RD_only	0.02	156	A	0.18	0.08	0.45	5.0x10 ⁻⁰⁷	0.19	0.08	0.46	5.8x10 ⁻⁰⁷
<i>RRM1</i>	rs116472045	A/G	11	0.02	3939	All	RD_only	0.04	157	H	2.64	1.42	4.90	0.0060	2.59	1.38	4.86	0.005081
<i>RTN3</i>	rs542998	T/C	11	0.43	3929	All	RD_only	0.41	147	H	0.88	0.63	1.22	0.4422	0.82	0.58	1.15	0.3977
<i>TRIM5</i>	rs7935564	G/A	11	0.44	3871	All	RD_only	0.47	151	R	1.40	0.95	2.05	0.0963	1.29	0.87	1.90	0.091014
<i>CAND1</i>	rs10459266	C/A	12	0.21	3938	All	RD_only	0.28	157	D	1.69	1.23	2.33	0.0014	1.75	1.26	2.42	0.0009
<i>CAND1</i>	rs12307123	C/T	12	0.23	3933	All	RD_only	0.26	156	A	1.22	0.94	1.58	0.1323	1.17	0.90	1.52	0.17144
<i>CAND1</i>	rs1566830	G/T	12	0.26	3940	All	RD_only	0.30	157	R	1.76	1.05	2.97	0.0449	1.76	1.04	2.98	0.0409
<i>GRIP1</i>	rs1394263	T/C	12	0.26	3914	All	RD_only	0.30	157	R	1.76	1.05	2.96	0.0459	1.74	1.03	2.95	0.041826
<i>IL22</i>	rs1012356	A/T	12	0.49	3940	All	RD_only	0.47	156	R	0.85	0.57	1.26	0.4147	0.88	0.59	1.31	0.4368
<i>IL22</i>	rs2227478	A/G	12	0.38	3939	All	RD_only	0.40	156	H	1.19	0.86	1.64	0.2908	1.25	0.90	1.73	0.24654
<i>IL22</i>	rs2227485	G/A	12	0.42	3928	All	RD_only	0.40	156	R	0.68	0.42	1.09	0.0939	0.67	0.41	1.08	0.0977
<i>IL22</i>	rs2227491	C/T	12	0.40	3870	All	RD_only	0.41	151	H	1.14	0.82	1.58	0.4365	1.13	0.81	1.57	0.38947
<i>IL22</i>	rs2227507	T/C	12	0.04	3948	All	RD_only	0.04	156	R	-	-	-	-	-	-	-	-
<i>TPTE2</i>	rs182873742	C/G	13	0.00	3940	All	RD_only	0.00	157	H	-	-	-	-	-	-	-	-
<i>RPS6KL1</i>	rs3742785	C/A	14	0.29	3714	All	RD_only	0.25	148	R	0.69	0.36	1.33	0.2424	0.83	0.63	1.08	0.2026
<i>SPTB</i>	rs229587	T/C	14	0.34	3872	All	RD_only	0.30	155	A	0.82	0.64	1.05	0.1067	0.82	0.64	1.05	0.10252
<i>YLP1</i>	rs10139016	C/T	14	0.25	3937	All	RD_only	0.24	157	R	0.78	0.38	1.60	0.4752	0.78	0.37	1.61	0.5046
<i>ADCY9</i>	rs10775349	C/G	16	0.20	3941	All	RD_only	0.21	151	R	1.75	0.90	3.40	0.1217	1.90	0.97	3.71	0.11955
<i>ADCY9</i>	rs2230739	A/G	16	0.09	3944	All	RD_only	0.10	151	R	2.58	0.78	8.59	0.1706	3.13	0.92	10.67	0.1687
<i>HBA</i>	α^{3-7} -thalassaemia	I/D	16	0.41	3946	All	RD_only	0.37	153	D	0.72	0.52	0.98	0.0340	0.76	0.54	0.98	0.036851
<i>IL4</i>	rs2243250	T/C	16	0.20	3902	All	RD_only	0.18	154	R	0.45	0.14	1.42	0.1209	0.47	0.15	1.51	0.1230
<i>ILAR</i>	rs1805015	T/C	16	0.42	3927	All	RD_only	0.44	156	D	1.10	0.78	1.55	0.5977	1.12	0.81	1.56	0.63147
<i>ADORA2B</i>	rs2535611	T/C	17	0.10	3911	All	RD_only	0.13	151	D	1.26	0.86	1.86	0.2420	1.28	0.86	1.89	0.2169
<i>NOS2</i>	rs1800482	G/C	17	0.08	3922	All	RD_only	0.09	154	R	1.96	0.46	8.34	0.4061	2.50	0.56	11.15	0.40375
<i>NOS2</i>	rs2297518	G/A	17	0.14	3910	All	RD_only	0.14	149	R	2.08	0.89	4.87	0.1234	2.02	0.85	4.79	0.1207
<i>NOS2</i>	rs8078340	C/T	17	0.21	3948	All	RD_only	0.20	156	R	0.15	0.02	1.04	0.0061	0.13	0.02	0.97	0.006273
<i>NOS2</i>	rs9282799	C/T	17	0.03	3949	All	RD_only	0.02	155	A	0.63	0.25	1.54	0.2725	0.66	0.27	1.62	0.2840
<i>TBX2</i>	rs73991577	A/G	17	0.00	3937	All	RD_only	0.00	157	H	-	-	-	-	-	-	-	-
<i>EMR1</i>	rs373533	G/T	19	0.46	3863	All	RD_only	0.47	151	R	0.99	0.67	1.47	0.9646	1.04	0.72	1.51	0.9102
<i>EMR1</i>	rs461645	C/T	19	0.47	3933	All	RD_only	0.47	151	D	1.02	0.71	1.47	0.9061	1.07	0.73	1.55	0.80482
<i>ICAM</i>	rs5498	A/G	19	0.11	3908	All	RD_only	0.10	154	R	0.44	0.06	3.19	0.3493	0.44	0.06	3.22	0.3534
<i>GNAS</i>	rs8386	C/T	20	0.19	3944	All	RD_only	0.18	155	H	0.89	0.63	1.27	0.5190	0.90	0.63	1.29	0.52994
<i>DERL3</i>	rs1128127	G/A	22	0.50	3911	All	RD_only	0.47	156	D	0.87	0.61	1.23	0.4269	0.88	0.61	1.26	0.3972
<i>CD40LG</i>	rs1126535	T/C	X	0.15	3934	M+F	RD_only	0.10	156	D	0.60	0.38	0.94	0.0184	0.59	0.37	0.94	0.020226
<i>CD40LG</i>	rs1126535	T/C	X	0.16	1947	F	RD_only	0.10	80	A	0.59	0.35	0.99	0.0311	0.61	0.37	1.00	0.0351
<i>CD40LG</i>	rs1126535	C/T	X	0.85	1987	M	RD_only	0.89	76	HM	1.49	0.71	3.13	0.2740	1.50	0.73	3.17	0.2940
<i>CD40LG</i>	rs3092945	T/C	X	0.21	3927	M+F	RD_only	0.24	154	D	1.19	0.84	1.68	0.3412	1.15	0.81	1.64	0.3059
<i>CD40LG</i>	rs3092945	T/C	X	0.21	1942	F	RD_only	0.25	79	A	1.30	0.89	1.88	0.1786	1.32	0.90	1.89	0.1986
<i>CD40LG</i>	rs3092945	C/T	X	0.78	1985	M	RD_only	0.77	75	HM	0.99	0.57	1.71	0.9629	1.00	0.58	1.72	0.9786
<i>G6PD</i>	rs1050828	C/T	X	0.19	3940	M+F	RD_only	0.20	155	H	1.12	0.70	1.79	0.6468	1.05	0.64	1.70	0.64676
<i>G6PD</i>	rs1050828	C/T	X	0.20	1951	F	RD_only	0.22	79	A	1.12	0.75	1.67	0.5802	1.16	0.78	1.69	0.6102
<i>G6PD</i>	rs1050828	C/T	X	0.19	1989	M	RD_only	0.18	76	HM	0.98	0.54	1.77	0.9503	0.99	0.57	1.84	0.9808
<i>G6PD</i>	rs1050829	T/C	X	0.40	3928	M+F	RD_only	0.38	155	A	0.92	0.76	1.12	0.3931	0.79	0.51	1.21	0.4246
<i>G6PD</i>	rs1050829	T/C	X	0.41	1952	F	RD_only	0.41	79	R	0.96	0.52	1.77	0.9017	0.97	0.55	1.79	0.9617
<i>G6PD</i>	rs1050829	C/T	X	0.60	1976	M	RD_only	0.66	76	HM	1.27	0.78	2.05	0.3344	1.29	0.79	2.08	0.3784

* The rs8176719 polymorphism identifies blood-group O in the homozygous deletion form. The derived allele of rs8176746 identifies blood-group B. Chr = chromosome; DAF = derived allele frequency; MO = genotype model [A – additive, D – dominant, H – heterozygote, HM – hemizygous males, R – recessive] with respect to the derived allele; OR = Odds Ratio; LCI = Lower Confidence Interval (95%); UCI = Upper Confidence Interval; P = p-value using a logistic regression model as described in methods. § ORs and p-values unadjusted and adjusted for ethnic group, gender and *rs334*. Only those individuals solely having RD were included in this analysis.

Table S11: Odds ratios for RD only for all polymorphisms investigated.

Gene 1	Gene 2	No. of cases by genotype						No. of controls by genotype						P
		REF REF	REF HET	HET REF	HET HET	DER REF	DER HET	REF REF	REF HET	HET REF	HET HET	DER REF	DER HET	
HBA	HBB	820	16	994	22	250	13	1131	216	1649	288	537	92	0.02
HBA	ABO	412	369	462	457	129	110	725	501	1058	753	342	250	0.05
HBA	LPHN2	720	124	890	116	216	45	1147	195	1615	322	533	101	0.05
ARL14	HBB	1680	48	392	7	17	1	2471	464	424	70	31	3	0.06
INPP4B	RPS6KLI	938	826	89	64	0	1	1673	1229	250	184	8	3	0.07
HBA	ARL14	656	163	814	169	208	49	1026	166	1460	243	469	91	0.07
HBA	INPP4B	772	69	929	83	230	30	1188	1219	301	237	21	10	0.07
HBB	LOC727982	1844	278	44	13	7	4	2774	520	495	94	28	5	0.08
ABO	IL10	694	404	590	347	124	70	1239	728	839	582	156	87	0.08
HBB	LPHN2	1844	278	44	13	7	4	2774	520	495	94	28	5	0.09
LOC727982	RPS6KLI	524	426	435	396	71	70	1029	749	726	538	168	119	0.11
ATP2B4	FREM3	895	105	900	109	165	19	1425	288	1434	312	342	69	0.13
LOC727982	LPHN2	946	136	806	145	148	15	1764	330	1231	225	268	51	0.13
INPP4B	LOC727982	980	886	102	67	0	2	1835	1260	275	195	6	7	0.15
ATP2B4	RPS6KLI	451	430	474	409	102	54	849	628	871	640	208	147	0.20
ABO	INPP4B	928	102	901	69	183	13	1825	286	1308	190	221	32	0.20
ARL14	IL10	1014	648	239	145	14	4	1697	1061	295	171	16	16	0.23
ABO	ATP2B4	474	460	448	453	82	100	726	967	665	672	118	112	0.29
IL10	LPHN2	1128	163	690	117	82	16	1905	341	1185	224	203	54	0.30
HBA	FREM3	750	86	892	116	235	23	1129	204	1575	355	510	110	0.35
IL10	INPP4B	1179	113	752	57	83	15	1954	302	1234	174	215	40	0.36
ARL14	INPP4B	1601	143	362	38	14	4	2573	370	420	77	30	4	0.40
IL10	RPS6KLI	598	533	382	317	48	41	1108	829	690	508	135	83	0.42
ATP2B4	HBB	971	26	983	25	175	6	1487	244	1466	287	354	62	0.44
FREM3	INPP4B	1910	48	102	131	2	2	3122	83	255	408	12	20	0.45
ARL14	LOC727982	855	763	203	166	8	9	1604	1083	261	197	16	15	0.50
ATP2B4	INPP4B	911	95	932	81	172	12	1510	219	1527	242	360	55	0.50
HBA	IL10	489	318	613	368	156	92	791	473	1112	712	366	232	0.50
ARL14	FREM3	1554	183	357	42	13	4	2432	488	392	102	18	4	0.50
ABO	HBB	1006	26	957	25	192	5	1779	328	1282	210	205	51	0.51
ARL14	ATP2B4	802	796	182	190	6	10	1329	1318	202	244	14	17	0.54
IL10	LOC727982	640	565	400	346	45	40	1214	851	777	514	128	101	0.54
INPP4B	LPHN2	1736	271	164	24	1	1	2859	522	421	93	40	4	0.56
FREM3	HBB	1899	48	220	8	5	0	2716	478	552	109	32	7	0.58
FREM3	LOC727982	965	853	115	99	3	3	1744	1181	357	262	19	16	0.65
HBB	IL10	1270	791	31	24	8	3	1902	1196	349	211	19	11	0.66
FREM3	RPS6KLI	914	806	112	82	1	4	1570	1177	331	233	26	8	0.68
HBA	RPS6KLI	391	338	479	398	121	112	639	498	985	683	311	234	0.69
ARL14	RPS6KLI	819	705	185	161	6	10	1429	1070	252	176	18	13	0.74
ARL14	LPHN2	1511	235	347	51	16	2	2467	471	428	67	26	8	0.74
HBB	INPP4B	1944	181	53	4	10	1	2880	421	503	87	24	13	0.75
ABO	RPS6KLI	488	415	443	399	95	77	1042	751	737	552	126	95	0.75
FREM3	IL10	1146	731	140	78	3	2	1840	1169	392	280	24	13	0.76
ATP2B4	LOC727982	488	438	502	440	91	79	929	634	967	645	221	170	0.76
LPHN2	RPS6KLI	900	761	126	130	3	3	1625	1175	294	236	12	9	0.86
ABO	LPHN2	886	147	842	123	169	26	1756	354	1276	218	217	34	0.86
ABO	FREM3	913	112	873	96	174	21	1735	355	1228	257	201	49	0.88
HBA	ATP2B4	375	392	476	455	116	120	588	605	876	874	276	294	0.91
FREM3	LPHN2	1692	263	302	30	5	1	2694	494	552	113	33	7	0.91
HBA	LOC727982	414	362	497	443	127	116	740	479	1046	744	336	242	0.95
ATP2B4	IL10	588	370	595	376	111	65	1005	620	1017	640	238	153	0.97
ABO	LOC727982	498	451	486	415	100	84	1140	792	815	554	141	92	0.98
HBB	RPS6KLI	998	858	23	25	5	5	1639	1173	280	238	16	11	0.99

REF: reference homozygote genotype; HET: Heterozygote genotype; DER: derived allele homozygote genotype. p-values were adjusted for ethnicity and gender.

Table S12: Pairwise interactions between the polymorphisms (1 per gene) at autosomal loci at which we found significant associations in our primary analyses using the likelihood ratio approach (Main Text Table 2).

Gene	SNP ID	genotype	Hb						MCV						RBC					
			N	G mean	LCL	UCL	P value	Regress P*	N	G mean	LCL	UCL	P value	Regress P*	N	G mean	LCL	UCL	P value	Regress P*
<i>ATP2B4</i>	rs1541255	AA	1010	60.58	59.03	62.17	NA		963	73.67	73.09	74.25	NA		966	2.64	2.57	2.72	NA	
<i>ATP2B4</i>	rs1541255	AG	1016	62.22	60.63	63.84	0.33	0.13	969	73.59	73.01	74.18	0.98	0.77	969	2.70	2.63	2.78	0.49	0.20
<i>ATP2B4</i>	rs1541255	GG	185	63.72	59.97	67.70	0.29		176	73.55	72.20	74.92	0.99		177	2.76	2.59	2.95	0.44	
<i>IL10</i>	rs1800890	AA	1313	61.33	59.95	62.74	NA		1264	73.81	73.30	74.33	NA		1264	2.67	2.60	2.73	NA	
<i>IL10</i>	rs1800890	AT	821	61.48	59.74	63.28	0.99	0.40	774	73.11	72.47	73.77	0.22	0.10	778	2.68	2.59	2.76	0.99	0.43
<i>IL10</i>	rs1800890	TT	99	58.69	54.02	63.76	0.57		92	73.69	71.81	75.61	0.99		92	2.63	2.40	2.88	0.94	
<i>LPHN2</i>	rs72933304	CC	1912	61.77	60.62	62.95	NA		1828	73.64	73.22	74.06	NA		1829	2.68	2.63	2.74	NA	
<i>LPHN2</i>	rs72933304	AC	296	60.74	57.90	63.72	0.80	0.65	278	73.50	72.43	74.60	0.97	0.09	281	2.67	2.54	2.82	0.99	0.43
<i>LPHN2</i>	rs72933304	AA	8	63.64	47.50	85.10	0.98		7	79.81	72.70	87.50	0.21		7	2.65	1.90	3.69	1.00	
<i>LOC727982</i>	rs1371478	CC	1088	62.43	60.89	64.01	NA		1035	74.03	73.47	74.59	NA		1037	2.70	2.63	2.78	NA	
<i>LOC727982</i>	rs1371478	CT	958	61.23	59.61	62.88	0.55	0.03	915	73.29	72.70	73.89	0.18	0.05	918	2.67	2.59	2.75	0.79	0.15
<i>LOC727982</i>	rs1371478	TT	163	58.61	54.94	62.52	0.17		156	72.93	71.52	74.38	0.35		156	2.60	2.42	2.79	0.56	
<i>ARL14</i>	rs75731597	AA	1754	62.00	60.79	63.23	NA		1674	73.68	73.24	74.12	NA		1676	2.69	2.63	2.75	NA	
<i>ARL14</i>	rs75731597	AC	401	61.21	58.74	63.78	0.85	0.19	384	73.15	72.24	74.07	0.57	0.006	385	2.70	2.59	2.83	0.98	0.02
<i>ARL14</i>	rs75731597	CC	18	56.54	46.57	68.66	0.62		16	80.60	75.81	85.69	0.01		17	2.14	1.73	2.64	0.09	
<i>FREM3</i>	rs186873296	AA	1967	61.36	60.23	62.51	NA		1875	73.66	73.24	74.08	NA		1876	2.67	2.62	2.73	NA	
<i>FREM3</i>	rs186873296	AG	233	62.94	59.63	66.44	0.66	0.01	222	73.54	72.34	74.76	0.98	0.51	225	2.72	2.56	2.88	0.84	0.10
<i>FREM3</i>	rs186873296	GG	6	84.39	60.20	118.10	0.15		6	71.46	64.60	78.90	0.82		6	3.08	2.15	4.41	0.72	
<i>INPP4B</i>	rs77389579	GG	2019	61.40	60.28	62.53	NA		1927	73.68	73.26	74.09	NA		1928	2.67	2.62	2.73	NA	
<i>INPP4B</i>	rs77389579	GT	188	63.77	60.05	67.73	0.46	0.16	178	73.25	71.91	74.61	0.83	0.07	180	2.74	2.57	2.93	0.73	0.18
<i>INPP4B</i>	rs77389579	TT	2	56.58	31.50	101.00	0.96		2	62.09	52.10	73.80	0.13		2	2.90	1.55	5.41	0.96	
<i>ABO</i>	rs8176719	Non-O	1178	57.49	55.95	59.07	NA		1125	71.56	70.97	72.15	NA		1126	2.54	2.29	2.44	NA	
<i>ABO</i>	rs8176719	O	1045	62.87	61.29	64.49	0.80	0.35	996	73.78	73.20	74.36	0.13	0.08	999	2.76	2.68	2.84	0.96	0.45
<i>ABO</i>	rs8176746	CC	1529	6.19	6.06	6.32	NA		189	72.36	71.07	73.67	NA		1458	2.70	2.64	2.76	NA	
<i>ABO</i>	rs8176746	AC	622	6.02	5.83	6.23	0.35	0.08	942	73.56	72.97	74.15	0.23	0.23	598	2.61	2.52	2.70	0.26	0.07
<i>ABO</i>	rs8176746	AA	76	5.79	5.27	6.37	0.37		996	73.78	73.20	74.36	0.13		73	2.59	2.34	2.88	0.74	
<i>HBB</i>	rs334	AA	2159	61.27	60.20	62.35	NA		2058	73.55	73.15	73.94	NA		2062	2.67	2.62	2.72	NA	
<i>HBB</i>	rs334	AT	57	73.54	66.01	81.92	0.003	5.1x10 ⁻⁷	55	70.89	68.59	73.28	0.08	1.0x10 ⁻⁹	55	3.40	3.03	3.82	0.0002	8.57x10 ⁻¹¹
<i>HBB</i>	rs334	TT	11	30.07	23.52	38.45	5.1x10 ⁻⁸		11	89.56	83.18	96.44	6.1x10 ⁻⁷		11	1.13	0.87	1.46	4.3x10 ⁻¹⁰	
<i>RPS6KLI</i>	rs3742785	CC	1032	62.86	61.26	64.50	NA		985	73.54	72.97	74.12	NA		987	2.71	2.64	2.79	NA	
<i>RPS6KLI</i>	rs3742785	CA	896	59.98	58.35	61.67	0.04	0.001	855	73.73	73.12	74.36	0.90	0.24	857	2.63	2.55	2.71	0.28	0.01
<i>RPS6KLI</i>	rs3742785	AA	187	57.98	54.57	61.60	0.04		176	74.34	72.97	75.73	0.55		176	2.50	2.34	2.67	0.06	
α ^{-3.7} thalassemia		Norm	853	61.12	59.43	62.87	NA		819	76.78	76.16	77.41	NA		819	2.54	2.46	2.61	NA	
α ^{-3.7} thalassemia		Het	1026	62.23	60.66	63.85	0.62		974	72.88	72.34	73.43	5.4x10 ⁻¹¹	8.3x10 ⁻⁶¹	975	2.72	2.65	2.80	0.002	
α ^{-3.7} thalassemia		Hom	264	62.16	59.09	65.38	0.84	0.49	252	66.80	65.82	67.78	5.4x10 ⁻¹¹		254	3.03	2.87	3.20	6.6x10 ⁻⁸	1.11x10 ⁻¹⁰
<i>CD40LG</i> [§]	rs3092945	TT	1459	61.93	60.61	63.28	NA		1390	73.48	73.00	73.97	NA		1392	2.70	2.64	2.76	NA	
<i>CD40LG</i>	rs3092945	TC	388	60.00	57.55	62.56	0.38	0.12	366	74.66	73.70	75.63	0.08	0.28	369	2.56	2.45	2.68	0.12	0.28
<i>CD40LG</i> [§]	rs3092945	CC	379	60.52	58.02	63.13	0.61		367	72.75	71.82	73.70	0.37		366	2.68	2.56	2.81	0.97	
<i>G6PD</i> [§]	rs1050828	CC	1643	61.93	60.68	63.20	NA		1574	73.17	72.71	73.62	NA		1575	2.71	2.65	2.77	NA	
<i>G6PD</i>	rs1050828	CT	306	61.76	58.92	64.74	0.99	0.001	286	75.12	74.03	76.23	0.003	0.003	288	2.58	2.45	2.72	0.20	0.01
<i>G6PD</i> [§]	rs1050828	TT	271	57.04	54.26	59.97	0.008		257	74.33	73.19	75.48	0.15		258	2.53	2.39	2.67	0.06	

Only those polymorphisms with the most significant associations with severe malaria phenotypes (main text Table 2) were analysed with red blood cell indices.

N = number of individuals; G mean = geometric mean; LCL = lower confidence level (95%), UCL = upper confidence level (95%); HB = haemoglobin (g/l); MCV = mean cell volume (fL); RBC = red blood cell count ($\times 10^{12}/l$); P value; Regress P = adjusted for gender, age, rs334 and ethnicity using linear regression. [§]Hemizygous males are coded as homozygotes.

Table S13: Association of Red Blood Cell indices for all severe malaria cases.

Gene	SNPID	*Individual loci						§Combined - 13 loci		
		Observed scale		Logit scale		Probit scale		Observed scale	Logit scale	Probit scale
		Case-control population	General population	Case-control population	General population	Case-control population	General population	General population	General population	General population
HBB	rs334	0.0390	0.0282	0.0361	0.0334	0.0361	0.0033	0.0295	0.0354	0.0354
FREM3	rs186873296	0.0098	0.0071	0.0080	0.0078	0.0080	0.0334	0.0064	0.0076	0.0076
FREM3	rs186790584	0.0092	0.0066	0.0074	0.0073	0.0074	0.0078			
FREM3	rs149914432	0.0089	0.0064	0.0072	0.0071	0.0072	0.0073			
ABO	rs8176746	0.0065	0.0047	0.0049	0.0048	0.0049	0.0071	0.0041	0.0044	0.0045
ABO	rs8176719	0.0061	0.0044	0.0046	0.0046	0.0046	0.0048	0.0011	0.0012	0.0011
INPP4B	rs77389579	0.0055	0.0040	0.0044	0.0044	0.0044	0.0046	0.0001	0.0003	0.0003
HBA	$\alpha^{-3.7}$ -thalassaemia	0.0042	0.0031	0.0033	0.0033	0.0033	0.0044	0.0021	0.0023	0.0022
INPP4B	rs13103597	0.0034	0.0025	0.0026	0.0026	0.0026	0.0026			
LOC727982	rs1371478	0.0033	0.0024	0.0025	0.0025	0.0025	0.0025	0.0027	0.0030	0.0029
LOC727982	rs1371474	0.0031	0.0023	0.0024	0.0024	0.0024	0.0024			
ARL14	rs75731597	0.0031	0.0022	0.0023	0.0023	0.0023	0.0023	0.0019	0.0021	0.0020
ARL14	rs74954675	0.0026	0.0019	0.0020	0.0019	0.0020	0.0019			
ARL14	rs76033371	0.0025	0.0018	0.0019	0.0018	0.0019	0.0018			
IL10	rs1800890	0.0018	0.0013	0.0015	0.0015	0.0015	0.0015	0.0013	0.0015	0.0015
IL10	rs3024500	0.0018	0.0013	0.0013	0.0013	0.0013	0.0013			
RPS6KL1	rs3742785	0.0016	0.0012	0.0012	0.0012	0.0012	0.0012	0.0008	0.0009	0.0009
EMR1	rs373533	0.0015	0.0011	0.0012	0.0012	0.0012	0.0012	0.0012	0.0013	0.0014
EMR1	rs461645	0.0014	0.0010	0.0011	0.0011	0.0011	0.0011			
LPHN2	rs72933304	0.0014	0.0010	0.0011	0.0011	0.0011	0.0011	0.0007	0.0008	0.0008
ATP2B4	rs55868763	0.0013	0.0010	0.0010	0.0010	0.0010	0.0010	0.0007	0.0007	0.0008
ATP2B4	rs1541255	0.0013	0.0010	0.0010	0.0010	0.0010	0.0010	Total	0.0525	0.0571
USP38	rs28459062	0.0013	0.0010	0.0010	0.0010	0.0010	0.0010			
IL1A	rs17561	0.0013	0.0010	0.0010	0.0010	0.0010	0.0010			
LPHN2	rs4650365	0.0012	0.0009	0.0009	0.0009	0.0009	0.0009			
ATP2B4	rs10900585	0.0012	0.0009	0.0009	0.0009	0.0009	0.0009			
DERL3	rs1128127	0.0012	0.0009	0.0009	0.0009	0.0009	0.0009			
ATP2B4	rs4951074	0.0011	0.0008	0.0009	0.0009	0.0009	0.0009			
TRIM5	rs7935564	0.0011	0.0008	0.0008	0.0008	0.0008	0.0008			
LOC727982	rs10188961	0.0010	0.0007	0.0008	0.0008	0.0008	0.0008			
IL20RA	rs1555498	0.0010	0.0007	0.0008	0.0008	0.0008	0.0008			
GBP7	rs1803632	0.0010	0.0007	0.0008	0.0008	0.0008	0.0008			
IL22	rs2227485	0.0010	0.0007	0.0008	0.0008	0.0008	0.0008			
TLR4	rs4986790	0.0007	0.0005	0.0007	0.0007	0.0007	0.0007			
IRF1	rs2706384	0.0009	0.0007	0.0007	0.0007	0.0007	0.0007			
GRIP1	rs1394263	0.0010	0.0007	0.0007	0.0007	0.0007	0.0007			
LPHN2	rs72933350	0.0009	0.0007	0.0007	0.0007	0.0007	0.0007			
IL4	rs2243250	0.0009	0.0007	0.0007	0.0007	0.0007	0.0007			
HSPA1B	rs6457452	0.0009	0.0006	0.0007	0.0007	0.0007	0.0007			
ADCY9	rs2230739	0.0008	0.0006	0.0006	0.0006	0.0006	0.0006			
LPHN2	rs72933310	0.0008	0.0006	0.0006	0.0006	0.0006	0.0006			
CAND1	rs1566830	0.0008	0.0006	0.0006	0.0006	0.0006	0.0006			
IL22	rs2227491	0.0008	0.0006	0.0006	0.0006	0.0006	0.0006			
IL22	rs1012356	0.0008	0.0006	0.0006	0.0006	0.0006	0.0006			
PLEKHG1	rs141555199	0.0007	0.0005	0.0006	0.0006	0.0006	0.0006			
ADORA2B	rs2535611	0.0007	0.0005	0.0005	0.0005	0.0005	0.0005			
GNAS	rs8386	0.0007	0.0005	0.0005	0.0005	0.0005	0.0005			
PLEKHG1	rs55958968	0.0007	0.0005	0.0005	0.0005	0.0005	0.0005			
CAND1	rs10459266	0.0007	0.0005	0.0005	0.0005	0.0005	0.0005			
IL1B	rs1143634	0.0006	0.0004	0.0005	0.0005	0.0005	0.0005			
TBX2	rs73991577	0.0005	0.0004	0.0005	0.0005	0.0005	0.0005			
TLR1	rs4833095	0.0006	0.0004	0.0004	0.0004	0.0004	0.0004			
IL10	rs1800896	0.0005	0.0004	0.0004	0.0004	0.0004	0.0004			
NOS2	rs9282799	0.0005	0.0004	0.0004	0.0004	0.0004	0.0004			
AJAP1	rs6674631	0.0005	0.0004	0.0004	0.0004	0.0004	0.0004			
NOS2	rs8078340	0.0005	0.0004	0.0004	0.0004	0.0004	0.0004			
CR1	rs17047660	0.0005	0.0004	0.0004	0.0004	0.0004	0.0004			
ICAM	rs5498	0.0005	0.0003	0.0004	0.0004	0.0004	0.0004			
TPTE2	rs182873742	0.0004	0.0003	0.0003	0.0003	0.0003	0.0003			
TNF	rs1800629	0.0004	0.0003	0.0003	0.0003	0.0003	0.0003			
RTN3	rs542998	0.0004	0.0003	0.0003	0.0003	0.0003	0.0003			
IL17RD	rs6780995	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
CAND1	rs12307123	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
PLEKHG1	rs114726617	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
TLR6	rs5743810	0.0002	0.0001	0.0002	0.0002	0.0002	0.0002			
PLEKHG1	rs142712208	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
PLEKHG1	rs2131263	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
AJAP1	kgp15825649	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
MKI67	rs115947774	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
TNF	rs361525	0.0003	0.0002	0.0002	0.0002	0.0002	0.0002			
CTLA	rs2242665	0.0002	0.0002	0.0002	0.0002	0.0002	0.0002			
TNF	rs1800750	0.0002	0.0001	0.0002	0.0002	0.0002	0.0002			
TNF	rs3093662	0.0002	0.0002	0.0002	0.0002	0.0002	0.0002			
TLR6	rs5743809	0.0002	0.0002	0.0002	0.0002	0.0002	0.0002			
SDCI	rs11899121	0.0002	0.0002	0.0002	0.0002	0.0002	0.0002			
LTA	rs2239704	0.0002	0.0002	0.0002	0.0002	0.0002	0.0002			
IL13	rs20541	0.0002	0.0001	0.0002	0.0002	0.0002	0.0002			
TLR9	rs187084	0.0002	0.0001	0.0002	0.0002	0.0002	0.0002			
SNORD48	rs116288147	0.0002	0.0001	0.0001	0.0001	0.0001	0.0001			
ILAR	rs1805015	0.0002	0.0001	0.0001	0.0001	0.0001	0.0001			

<i>PLEKHG1</i>	rs15116938	0.0002	0.0001	0.0001	0.0001	0.0001	0.0001
<i>RRM1</i>	kgp12768002	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>CR1</i>	rs17047661	0.0002	0.0001	0.0001	0.0001	0.0001	0.0001
<i>SPTB</i>	rs229587	0.0002	0.0001	0.0001	0.0001	0.0001	0.0001
<i>PLEKHG1</i>	rs76924464	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>PLEKHG1</i>	rs151293197	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>PLEKHG1</i>	rs79100774	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>LTA</i>	rs909253	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>MKI67</i>	rs148494166	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>NOS2</i>	rs1800482	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>MKI67</i>	rs11016116	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>ABO</i>	rs8176750	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>HCG4</i>	rs114980857	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>TNF</i>	rs1799964	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>B3GALNT1</i>	rs12107243	0.0001	0.0001	0.0001	0.0001	0.0001	0.0001
<i>CD36</i>	rs3211938	0.0001	0.0000	0.0001	0.0001	0.0001	0.0001
<i>LPHN2</i>	rs146428334	0.0001	0.0000	0.0000	0.0000	0.0000	0.0000
<i>IL17RE</i>	rs708567	0.0001	0.0000	0.0000	0.0000	0.0000	0.0000
<i>NOS2</i>	rs2297518	0.0001	0.0000	0.0000	0.0000	0.0000	0.0000
<i>IL22</i>	rs2227478	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>ATP2B4</i>	rs3753036	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>NOD1</i>	rs2075820	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>ABO</i>	rs56390333	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>IL22</i>	rs2227507	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>GABBR1</i>	rs192151845	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>GAB1</i>	rs7663712	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>FREM3</i>	rs184895969	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>C6</i>	rs1801033	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>ADCY9</i>	rs10775349	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>FREM3</i>	rs184908374	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>DARC</i>	rs2814778	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000
<i>TLR4</i>	rs4986791	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

*Estimates are the proportion of the total variance explained by each polymorphism individually when estimated on the scale shown ("Observed" where computed on the 0-1 scale and transformed to the underlying liability scale; "Logit" and "Probit" where computed using the logit and probit transformations to model them on the liability scale). Estimates from the case-control data ("Case-control") are inflated due to ascertainment bias and thus are adjusted to give the values expected in the general population ("General"). Estimates are missing for markers that had <5 genotypes carrying the minor allele. The model here included sex as a fixed effect but not ethnic group. When ethnic group was included, the estimates decreased by a factor of 0.91 on average. §Estimates when simultaneously fitting the most significant SNP for each of 12 genes explaining the most amount of variance individually under the logit model. Two SNPs for the ABO locus are included because they independently explained significant amounts of variance. The total variance explained by these 13 markers is shown in bold below these with adjustment for ascertainment bias.

Table S14: Proportions of variance explained by individual markers

		BlpI RFLP assay for the Dantu hybrid gene		
		Non-Dantu	Dantu positive	Total
iPLEX assay for rs186873296	AA	1685*	13	1698
	AG	17	213*	230
	GG	1	9*	10
Total		1703	235	1938

All severe malaria cases (n=2244) were genotyped for the Dantu hybrid gene; data are shown for samples with both a valid rs186873296 and Dantu result. The BlpI RFLP assay does not distinguish readily between heterozygous and homozygous states for Dantu. RFLP=restriction fragment length polymorphism. *Concordant data between assays.

Table S15: Concordance between iPLEX assay for rs186873296 and the BlpI RFLP assay for the Dantu hybrid in paediatric cases of severe malaria

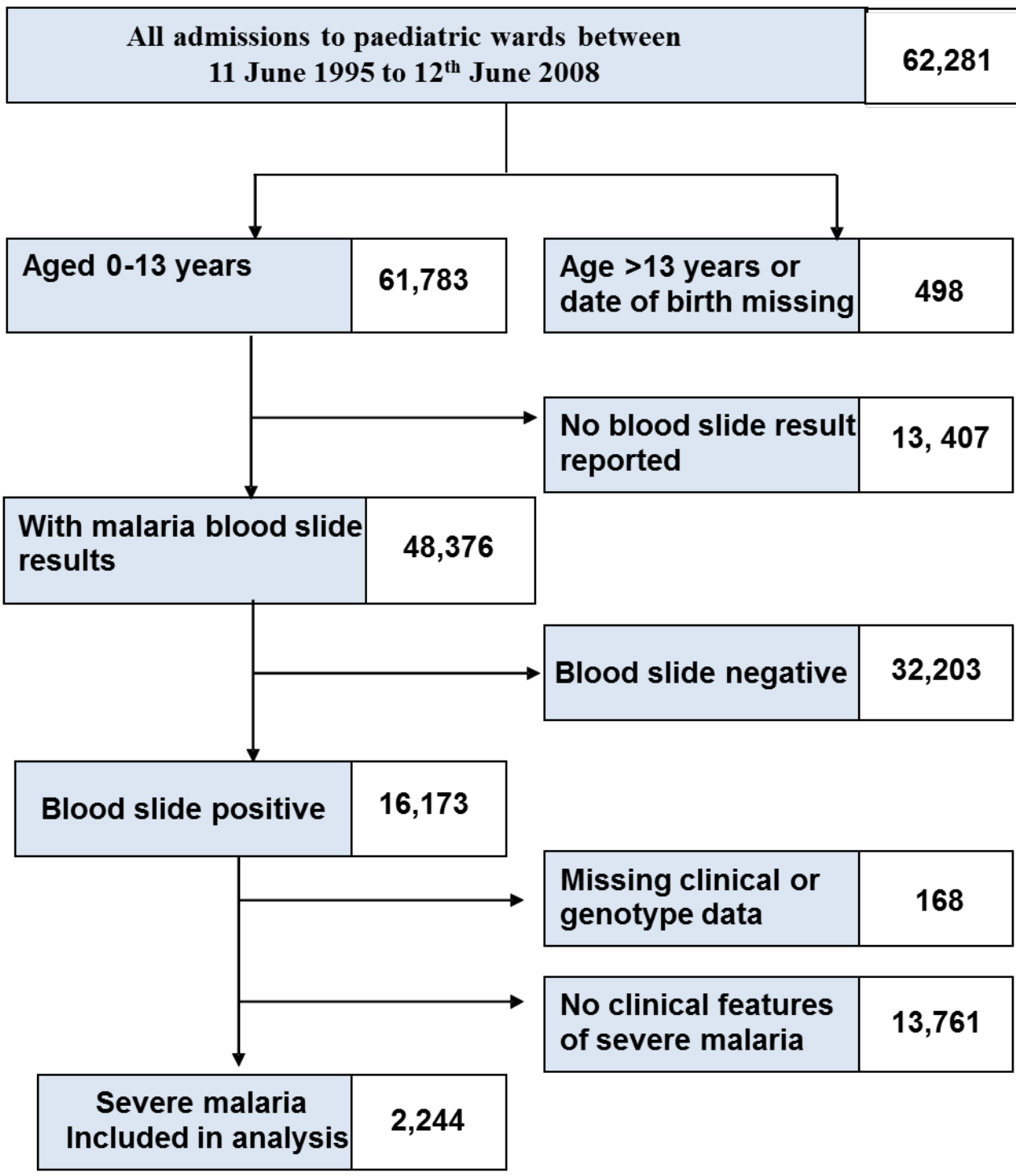


Figure S1: Flow diagram showing the selection of severe malaria cases.

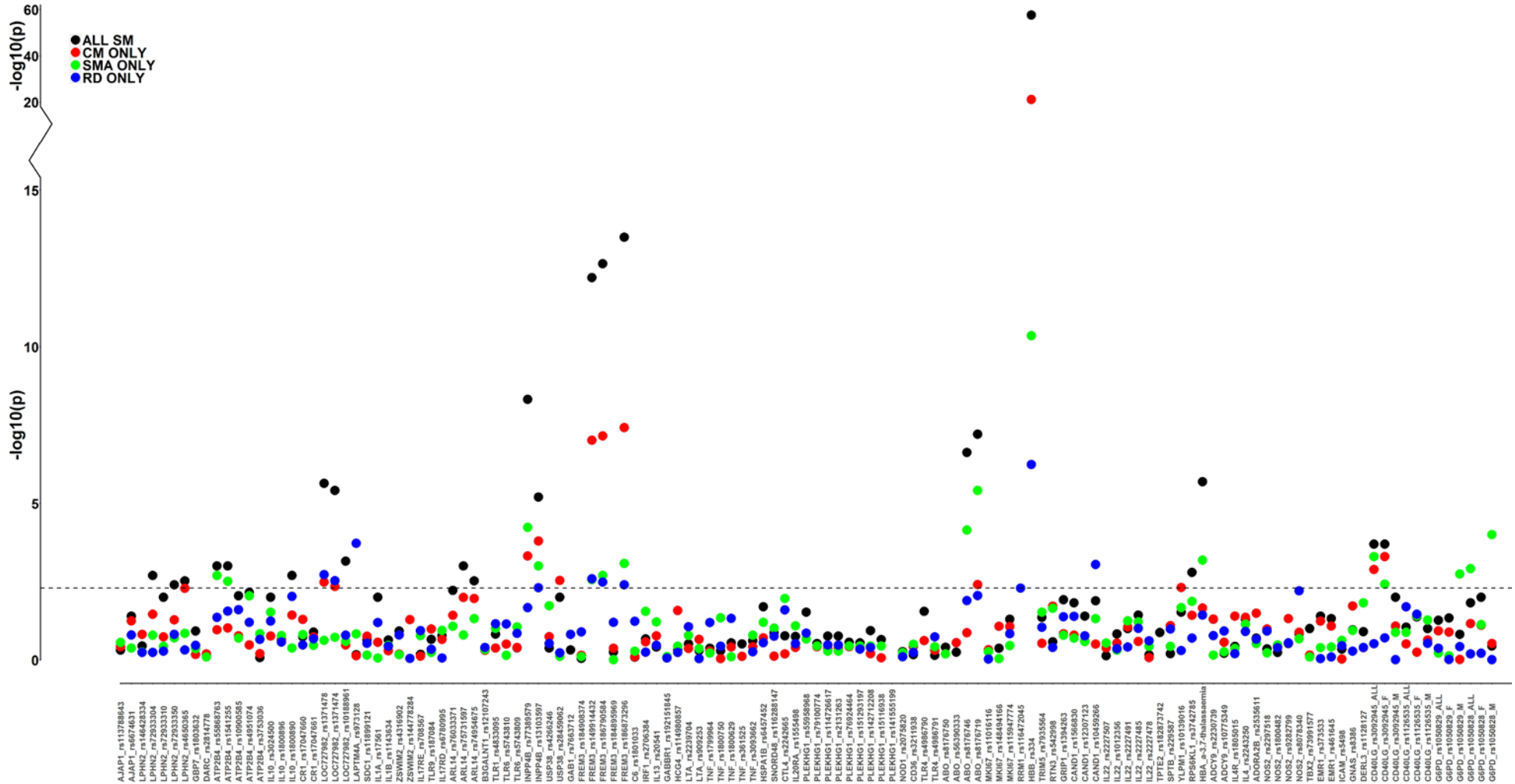
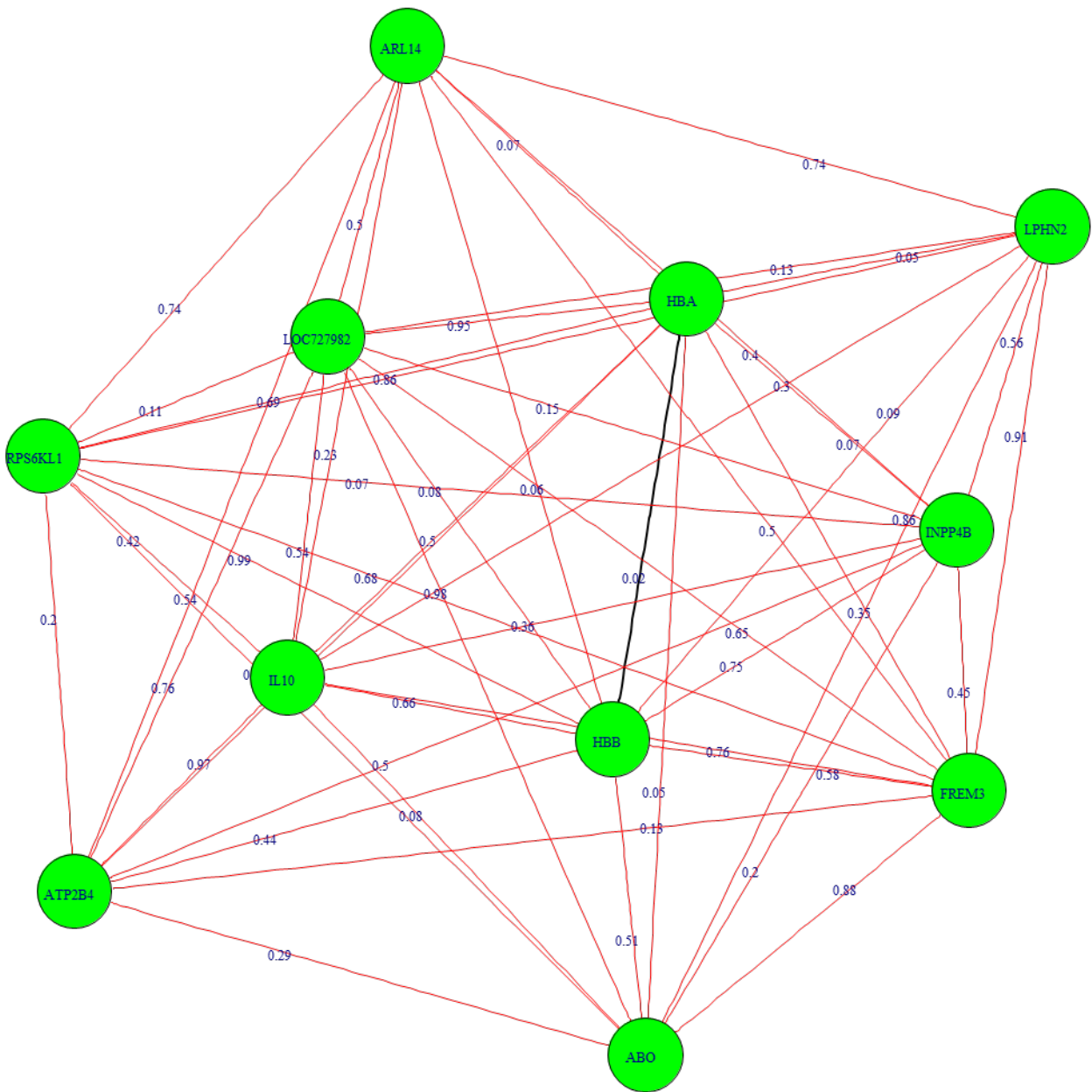


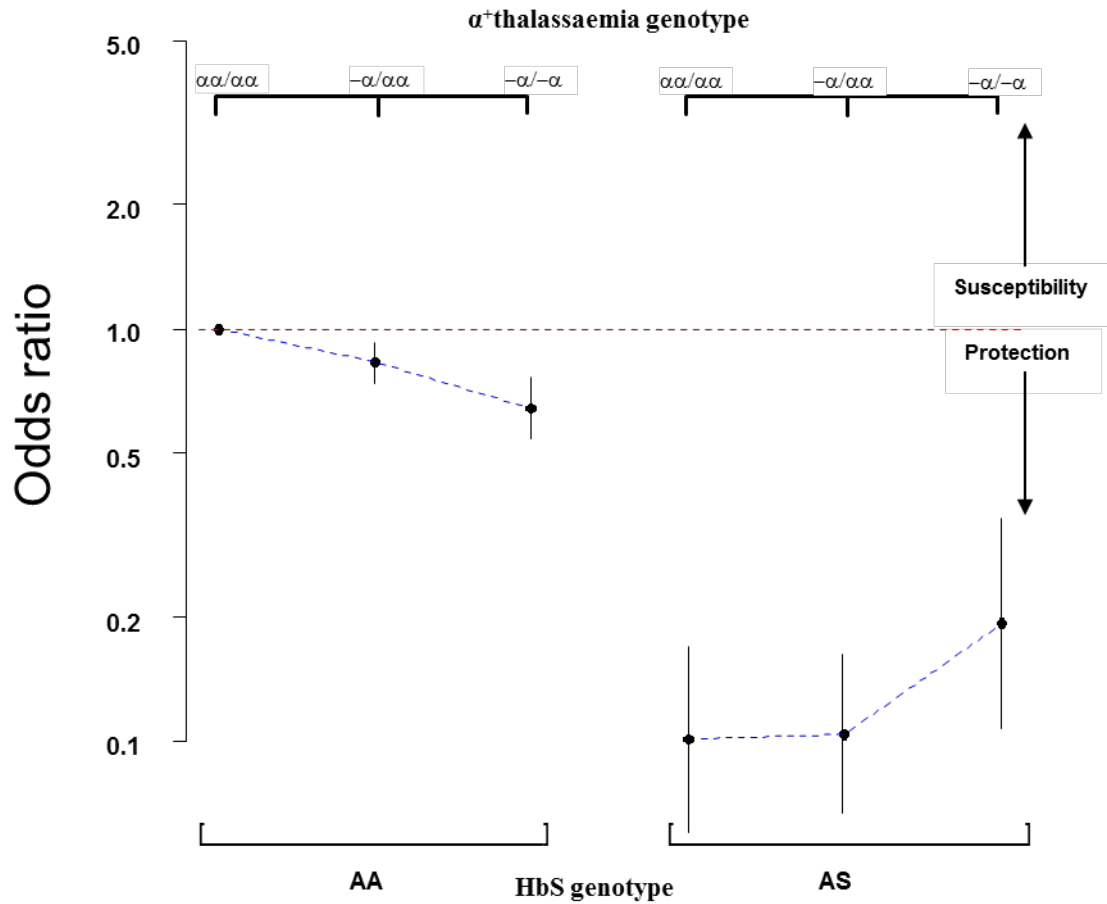
Figure S2. Manhattan plot showing the distribution of p-values for association tests with severe malaria phenotypes.

All SM; all individuals with any severe malaria phenotype; CM, cerebral malaria only; SMA, severe malaria anaemia only; RD, respiratory distress only. We tested for associations using additive (A), dominant (D), heterozygous (H), hemizygous (HM) and recessive (R) genetic models and the minimum p-value (as $-\log_{10}[p]$) for each polymorphism across the models is shown. Analyses were adjusted for gender, ethnic group and rs334 genotype. The dashed horizontal line represents the significance threshold of $p < 0.005$ determined by permutation testing (see methods). Polymorphisms tested are listed in chromosomal order (see Supplementary Table 1 for details).



The figure shows a graphical representation of the data in Supplementary Table S12. The polymorphisms used for the pairwise interaction analysis were those shown in the main text Table 2. The nodes in the figure are labelled according with the gene name for the polymorphisms. The lines connecting the nodes show the p-values for the interactions. p-values were adjusted for ethnicity and gender.

Figure S3: Network diagram for interaction of polymorphim pairs in severe malaria.

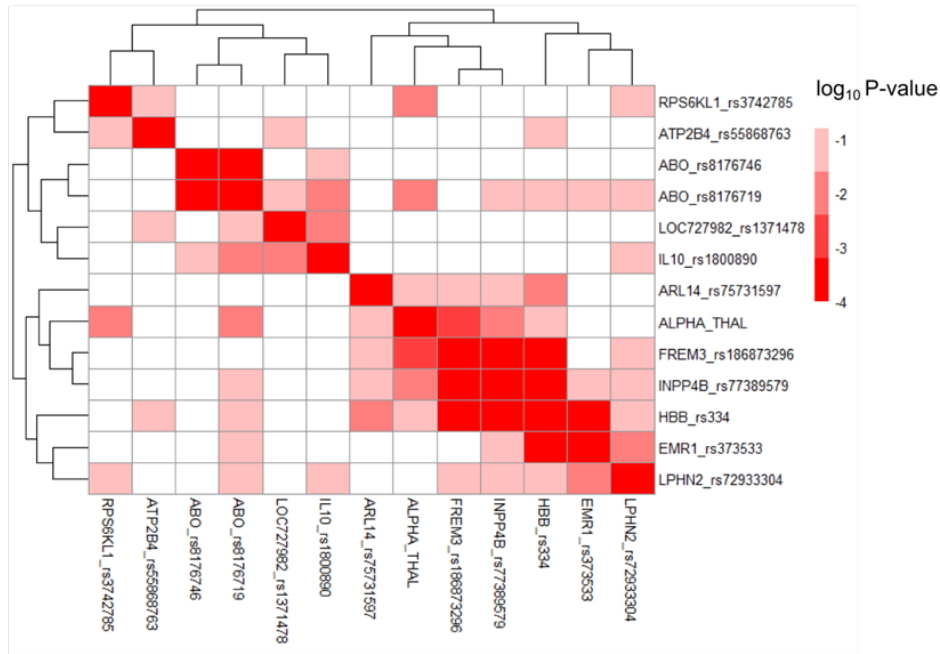


Allele group	a	b	c	d	e	f
Cases	828	1002	252	16	22	13
Controls	1131	1649	537	216	288	92
P-value	ref	0.002	5.8×10^{-7}	3.1×10^{-18}	1.4×10^{-23}	4.0×10^{-8}

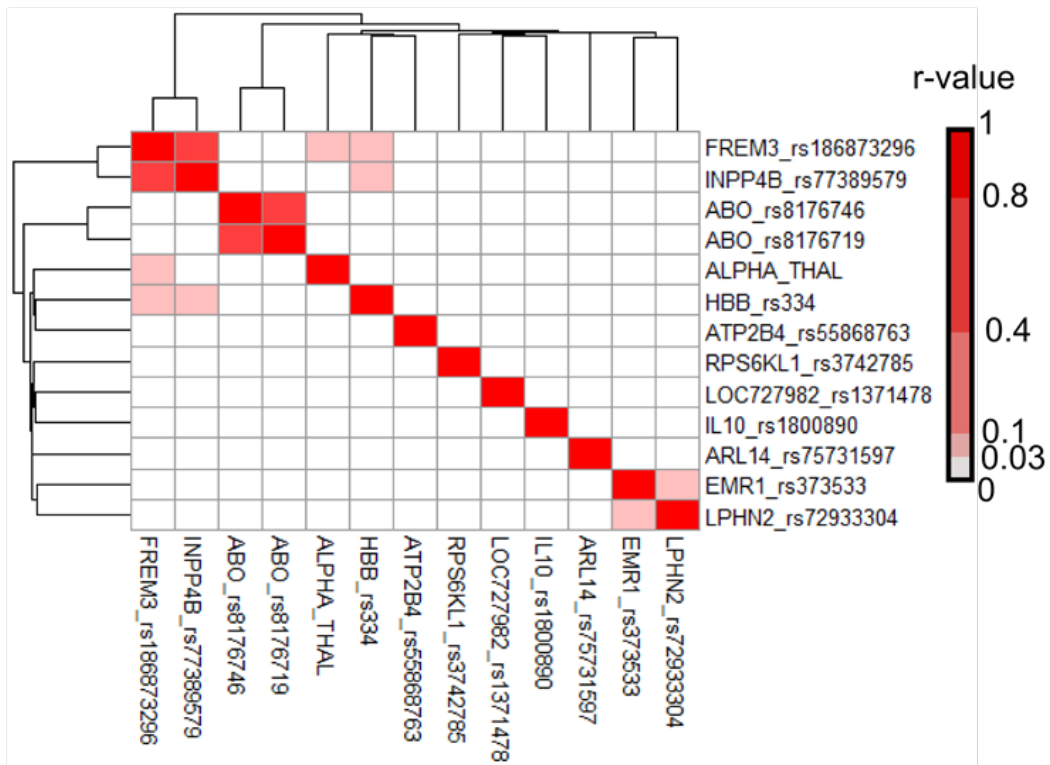
For each combination of genotypes (x-axis), we computed the odds ratio \pm 95% CI (y-axis) and p-value relative to the reference group (α^+ thalassaemia: $\alpha\alpha/\alpha\alpha/\text{HbAA}$). The red line shows the point of no effect. The odds ratio has been plotted on a log scale so that the distances above and below 1 represent the same size of effect although opposite effect. We also list the sample size for cases and controls. Allele group “a” shows the baseline for normal α^+ thalassaemia and HbAA. As shown in allele groups “a”, “b” and “c”, on a normal HbAA background, α^+ thalassaemia tends to protect reaching an OR of 0.6 in the homozygote state. However, this trend for protection is not seen in individuals with HbAS, indeed the α^+ thalassaemia locus reduces the effect of HbAS (OR 0.1 to 0.2).

Figure S4: Two-way epistatic interactions between rs334 (HbS) and $\alpha^{3.7}$ -thalassaemia for all severe malaria cases.

A



B



Panels A and B show the p-value and r-value for each pairing of the 13 polymorphisms (Figure 3 and Supplementary Table 14) explaining the variability in severe malaria among individuals in the general population (one SNP per gene except for ABO where both rs8176719 and 8176746 independently explained significant amounts of variance).

A: p-value: Colour indicates significance of linkage disequilibria by χ^2 test. polymorphism pairs are clustered and ordered according to closeness of p-values.

B: r-value: Colour indicates the strength of the correlation between genotypes (r-value). Values of $r > 0.03$ are highly significant by χ^2 test ($P < 0.001$). polymorphism pairs are clustered and ordered according to relatedness of r-values.

Figure S5. Linkage disequilibria among the 13 polymorphisms explaining most of the variance in the population.

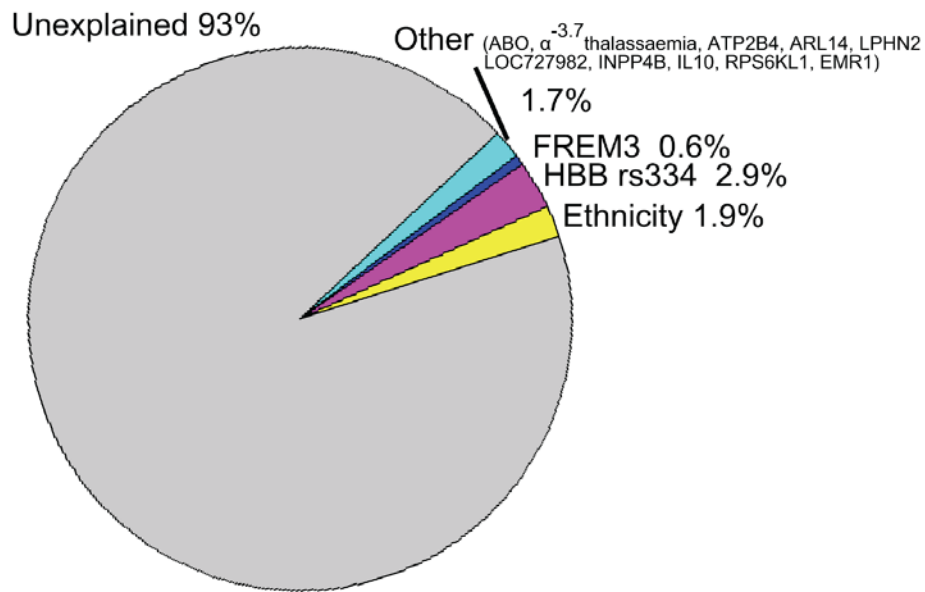


Figure S6. Proportion of variance in the risk of severe malaria among children that is explained by genetic polymorphisms and other factors. Values pertain to the general population and were obtained by analysing on the observed (0-1) scale and then transforming to the underlying liability scale (see Supplementary Table 12).

MalariaGEN Consortium members

This list can also be viewed at;

<https://www.malariagen.net/projects/consortial-project-1/malariagen-consortium-members>

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