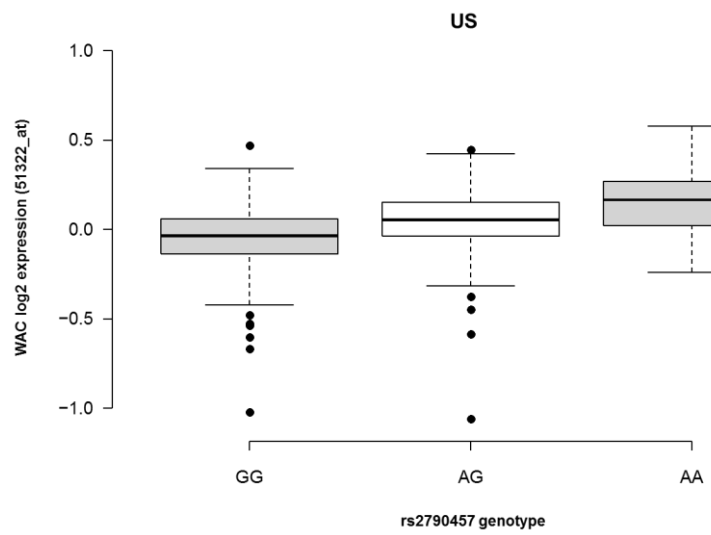
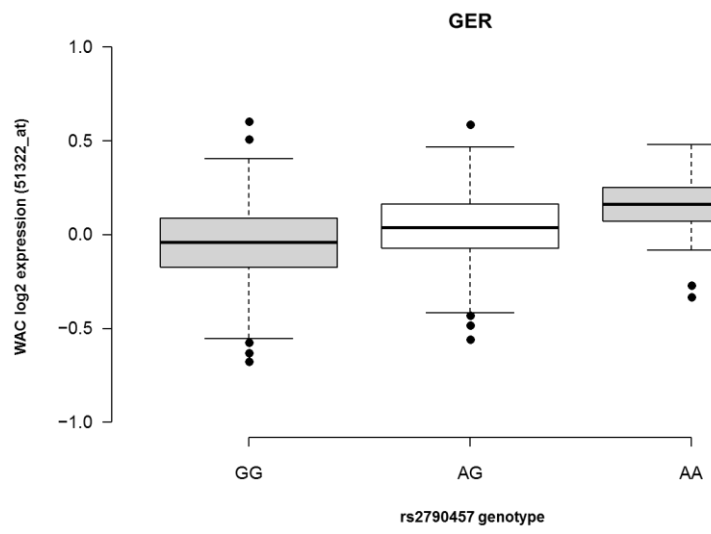
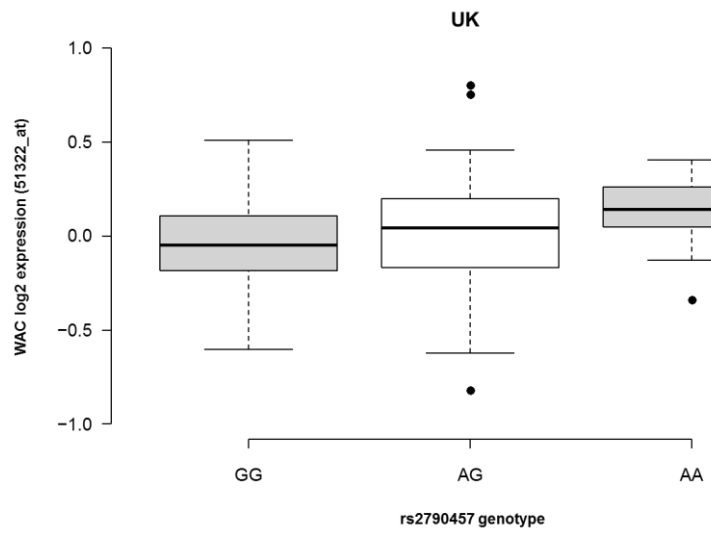
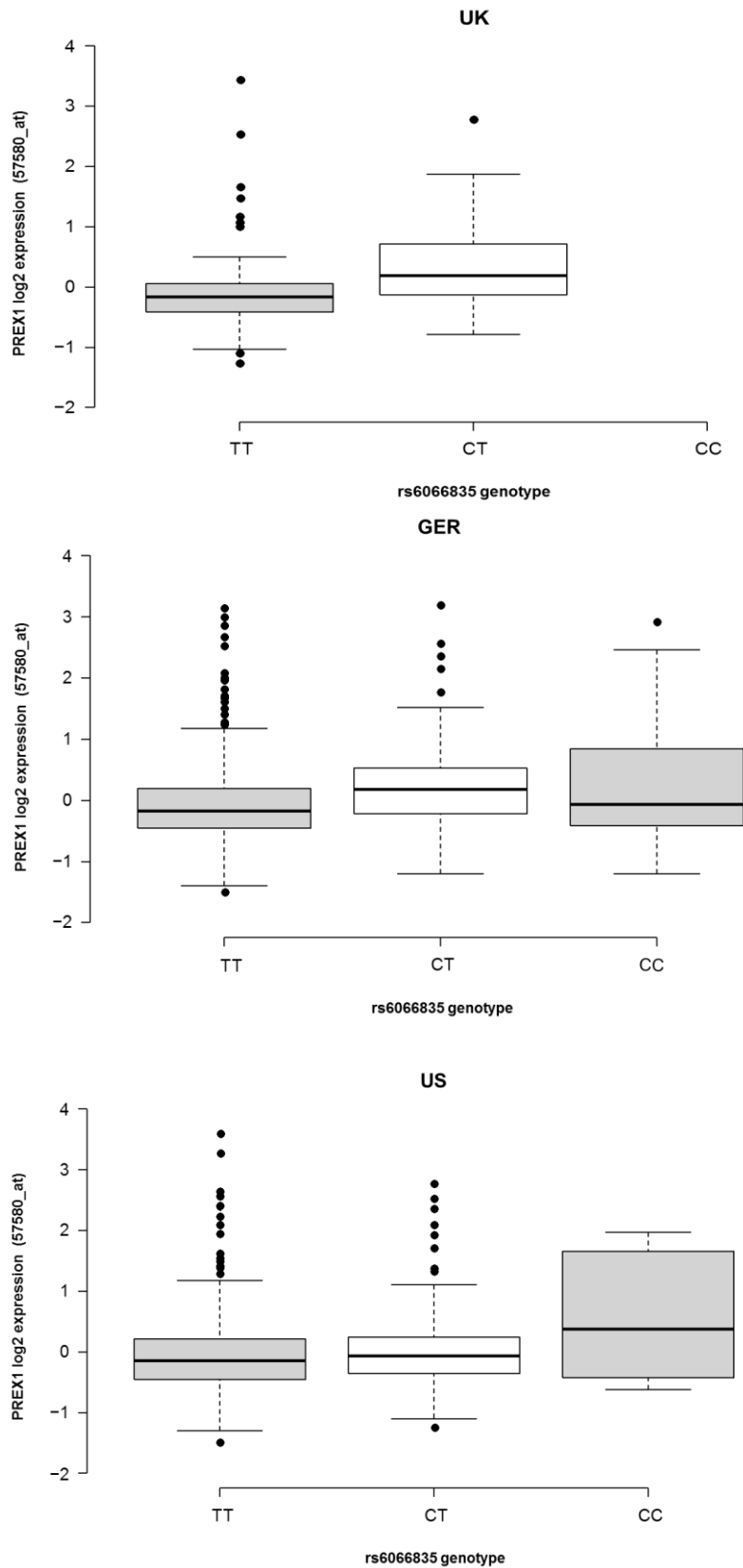
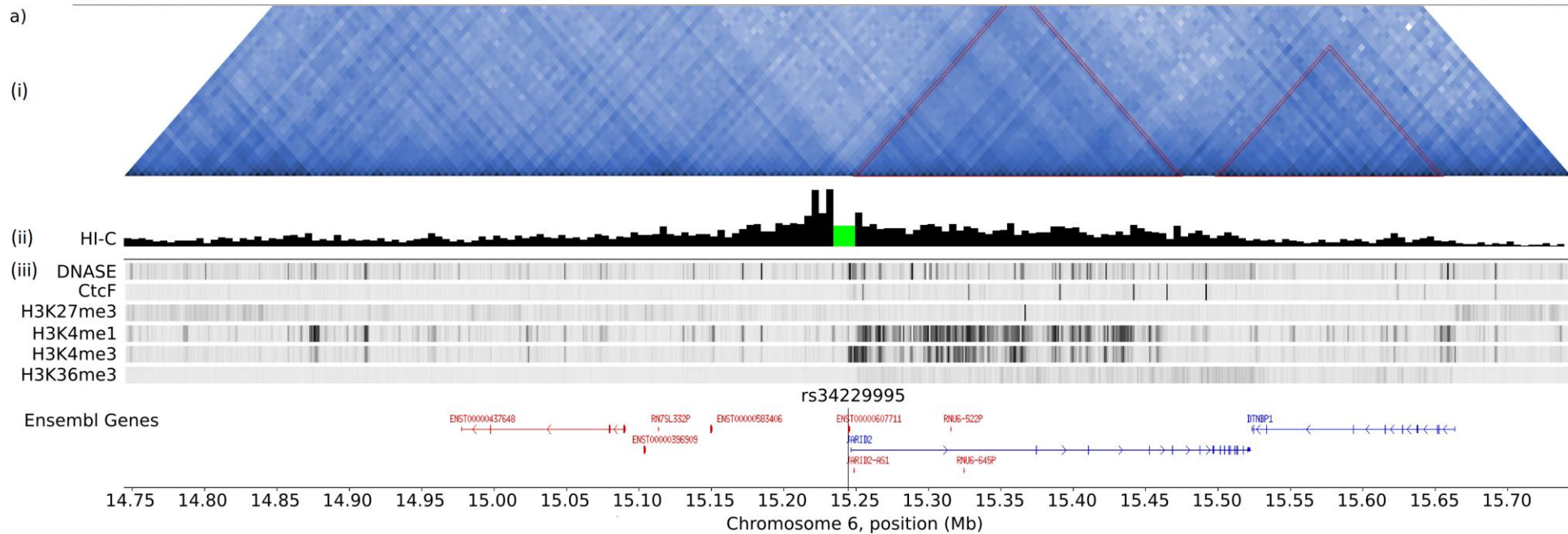


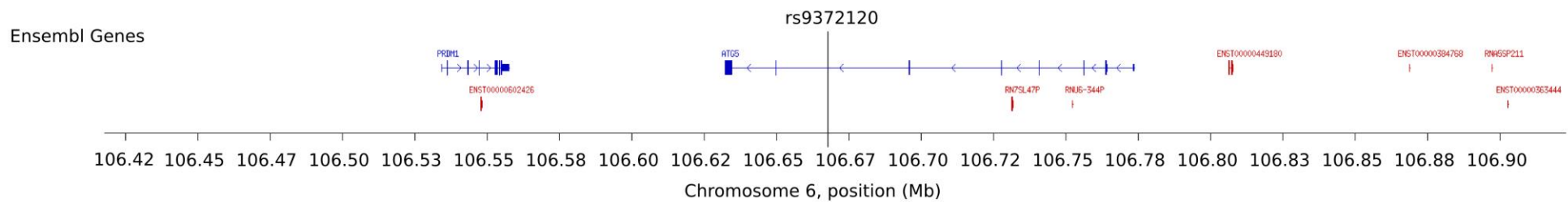
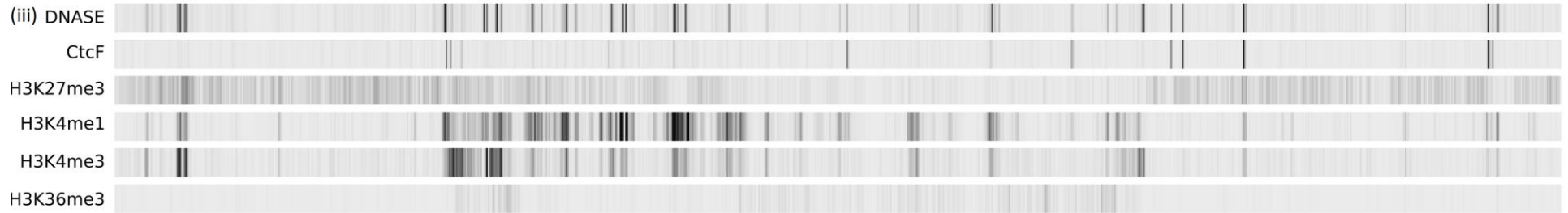
Supplementary Figure 1: Quantile-Quantile (Q-Q) plots of observed and expected χ^2 values of association between SNP genotype and risk of multiple myeloma after imputation. a) UK, b) Sweden/Norway, c) Germany, d) Iceland, e) USA and f) Netherlands. The blue line represents the null hypothesis of no true association.

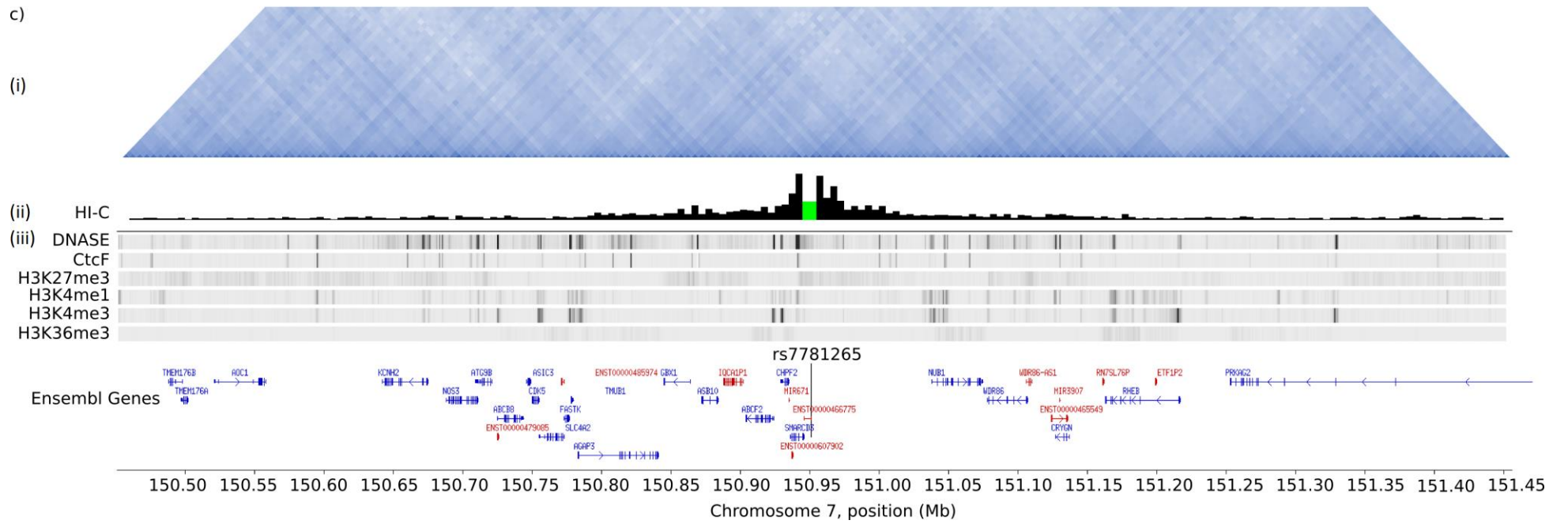


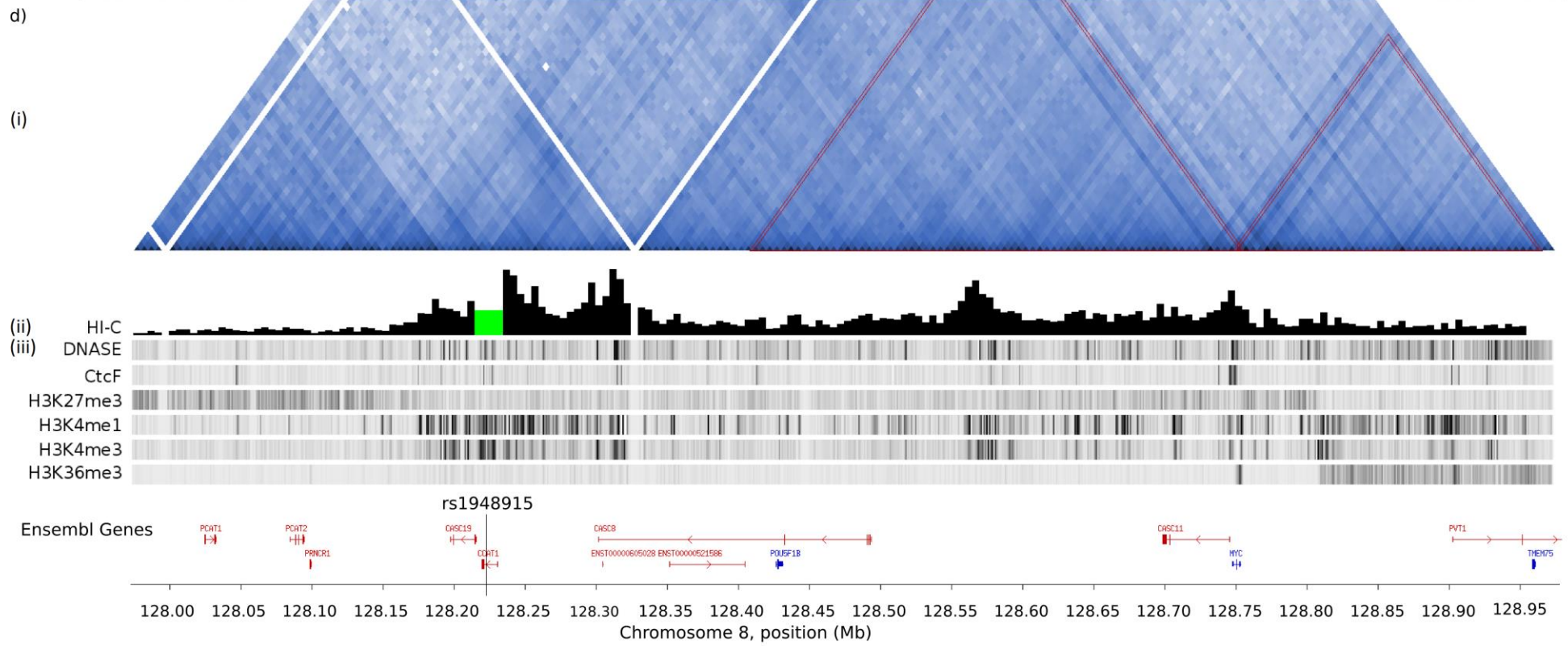


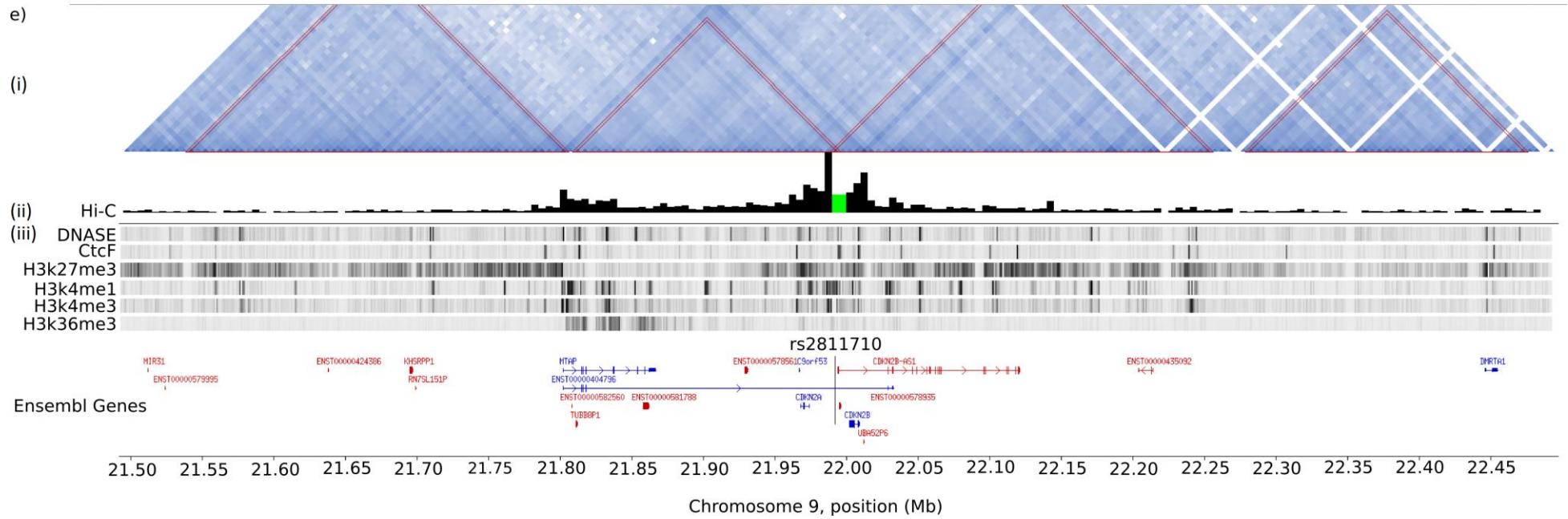
Supplementary Figure 2: eQTL boxplots. Relationship between rs2790457 (10p12.1) genotype and *WAC* expression, and rs6066835 (20q13.13) and *PREX1* in CD138+ selected plasma cells from 658 German, 183 UK and 608 US myeloma patients. The central line in each box indicates the median; the bottom and top lines of the box are the 25th and 75th percentiles; whiskers extend 1.5 times from the 25th and 75th percentiles.

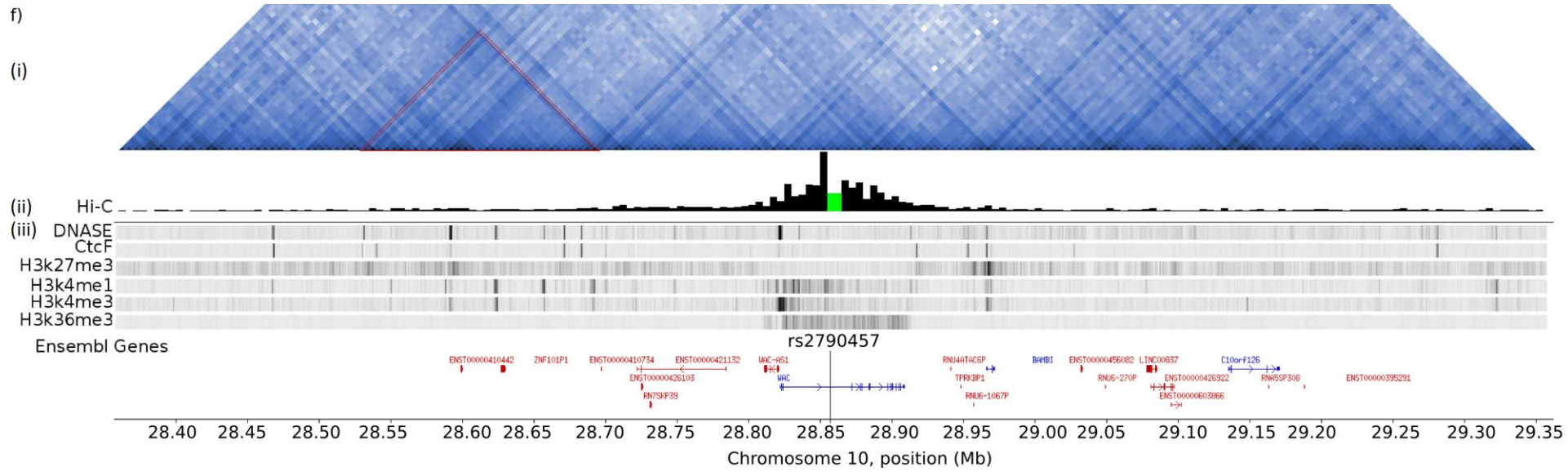


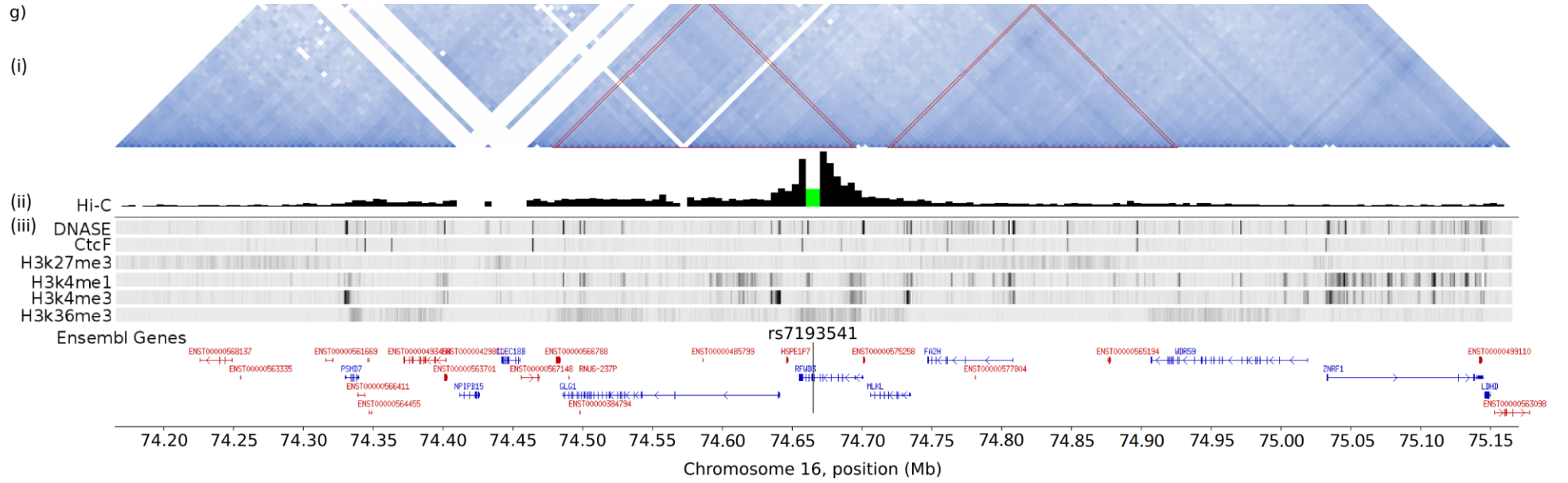


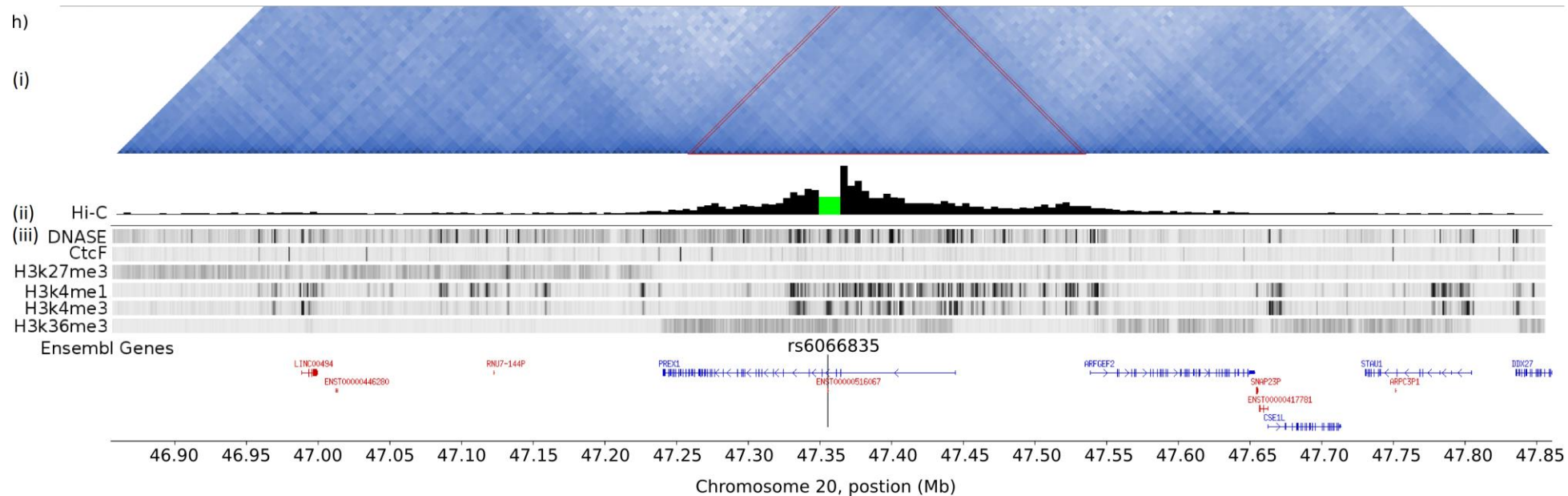






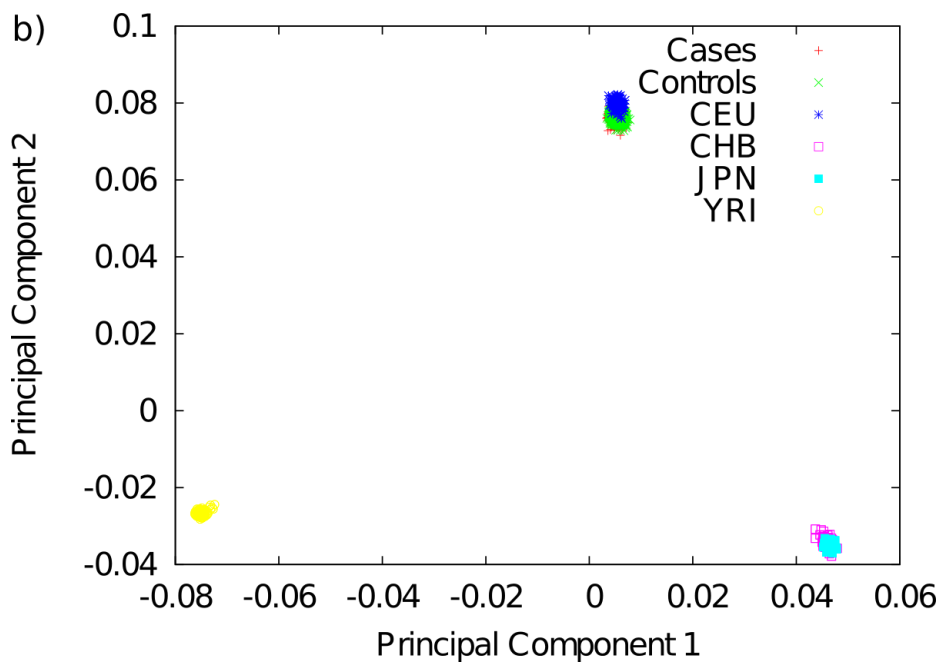
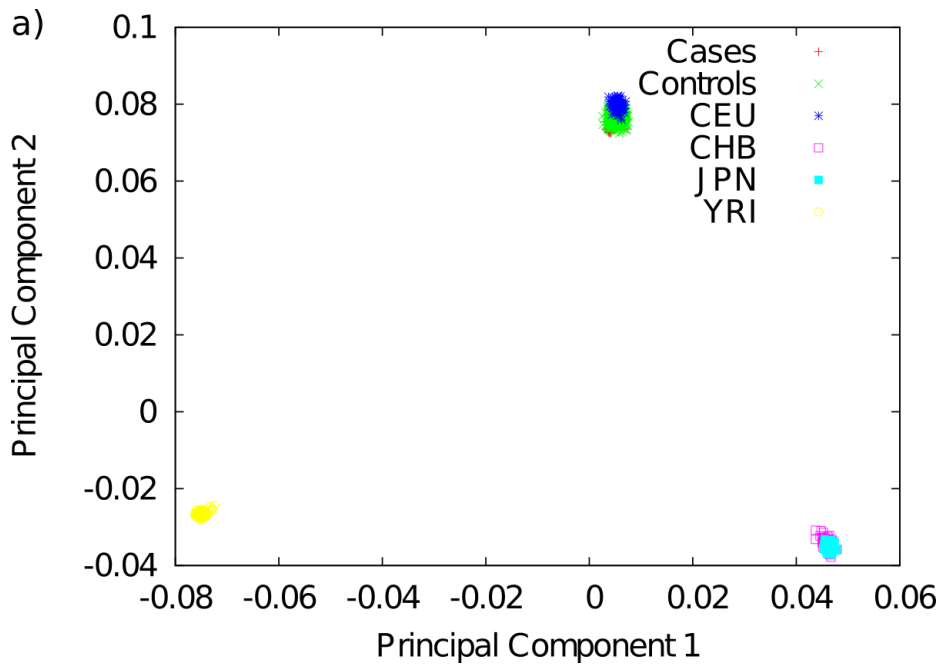


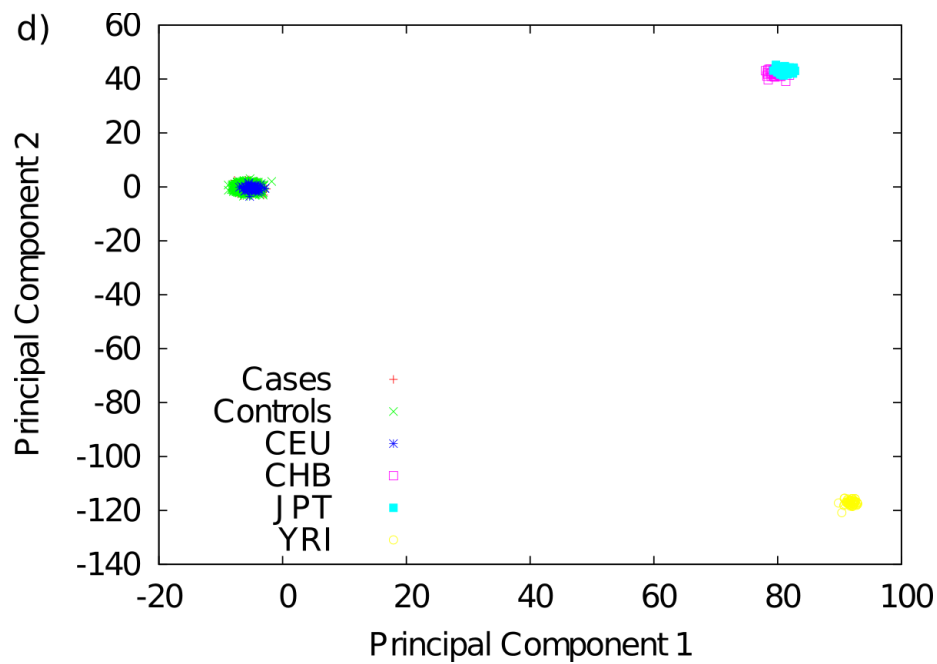
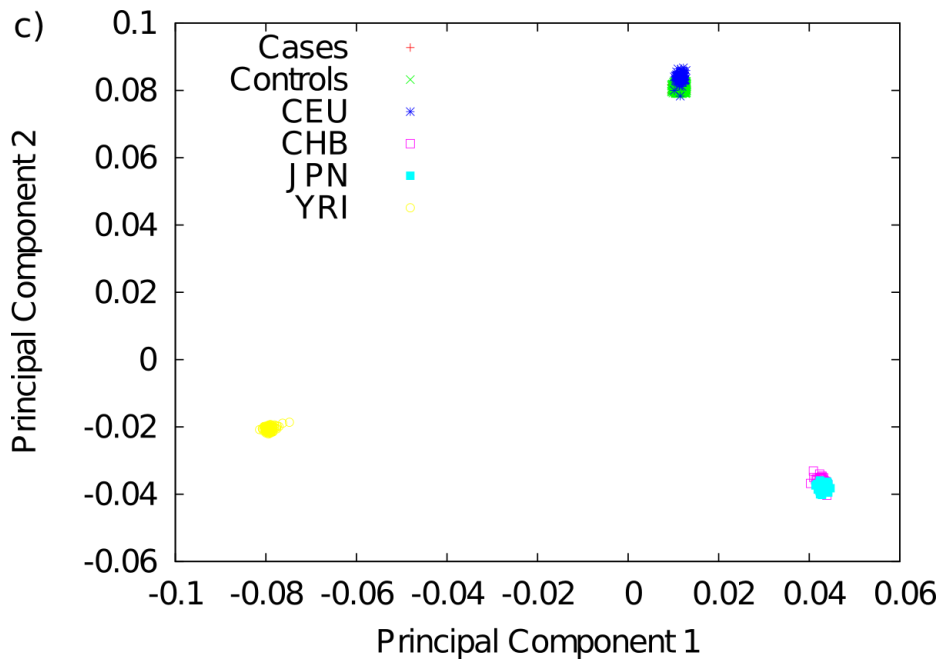




Supplementary Figure 3: Definition of topological domains (TADs) at risk loci and patterns of local chromatin interactions in GM12878¹.

(i) The upper panel shows heat map of chromatin interactions. TADs at each locus (double-red line) were inferred by the Arrowhead method and are superimposed. (ii) Middle panel denotes virtual 4C experiment defined by Hi-C intensity according to viewpoint based on sentinel SNP. The green box denotes a 5Kb (masked) region flanking the sentinel SNP. (iii) Lower panel shows DNase, CTCF, H3K27me3, H3K4me1, H3K4me3 and H3K36me3 tracks for GM12878.





Supplementary Figure 4: Identification of individuals of non-European ancestry in cases and controls.

(a) UK-GWAS (Thresholds; $0.0025 < PC1 < 0.0076$, $0.0724 < PC2 < 0.0804$), (b) German-GWAS (Thresholds; $0.0033 < PC1 < 0.0079$, $0.0716 < PC2 < 0.0803$), (c) USA-GWAS (Thresholds; $0.0096 < PC1 < 0.0129$, $0.0791 < PC2 < 0.0829$) and (d) Netherlands-GWAS (Thresholds; $-8.8 < PC1 < -1.7$, $-3.5 < PC2 < 3.1$). The first two principal components of the analysis are plotted. HapMap CEU individuals are plotted in blue; CHB individuals are plotted in indigo; JPN are plotted in cyan; YRI individuals are plotted in yellow. Cases are plotted in red, controls plotted in green, and have been removed if observed here to be of non-European ancestry.

	UK		Sweden/Norway		Germany		Netherlands		USA		Iceland	
	Cases	Controls	Cases	Controls	Cases	Controls	Cases	Controls	Cases	Controls	Cases	Controls
Pre-QC	2,329	5,199			1,512	2,107	608	2669	1,076	2,234		
Sex discrepancy	10	0			1	0	0	0	0	0		
Call rate fail	1	0			0	0	2	0	0	4		
Heterozygosity rate	NA	NA			NA	NA	7	0	9	2		
Related Individuals	2	2			0	0	0	0	1	0		
Non-European Ancestry	34	0			3	0	44	0	286	369		
Post-QC	2,282	5,197	1,714	10,391	1,508	2,107	555	2669	780	1,857	480	212,164

Supplementary Table 1: Details of the quality control filters applied to each GWAS. Samples were excluded due to call rate (<95% or failed genotyping), ethnicity (principle components analysis or other samples reported to be not of white, European descent), relatedness (any individuals found to be duplicated or related within or between data sets through IBS) or sex discrepancy. Sweden/Norway and Iceland: These studies have been previously reported in their entirety with full QC details².

	UK	Sweden/Norway	Germany	Netherlands	USA	Iceland
Pre-QC	409,429		401,405	646,124	296,998	
Call rate fail	997		113	6,523	4	
HWE fail	7		0	18,104	171	
MAF < 0.01	3		1	0	9151	
Post-QC	408,422		401,291	621,497	287,672	
Imputed (filtered)	8,517,071	7,182,761	8,282,831	8,628,799	8,085,846	10,291,845

Supplementary Table 2: Details of the quality control filters applied to each GWAS. Genotyped SNPs with a call rate <95% were excluded as were those with a MAF<0.01 or displaying significant deviation from Hardy-Weinberg equilibrium (*i.e.* $P < 10^{-5}$). Imputed SNPs with information score <0.8 and MAF <0.005 were excluded. Sweden/Norway and Iceland: These studies have been previously reported in their entirety with full QC details².

Locus	Published SNP	Strongest signal (current GWAS)	Position (hg19, bp)	Risk Allele	Literature OR (95% CI)	Literature <i>P</i> value	Current GWAS OR (95% CI)	Current GWAS <i>P</i> value	Reference																																																																																														
2p23.3	rs6746082		25659244	A	1.29 (1.17-1.42) $r^2 = 0.483$	1.22×10^{-7} $D' = 0.713$	1.20 (1.15-1.26) 1.24 (1.18-1.30)	2.28×10^{-14} 1.24×10^{-16}	3																																																																																														
		rs7577599	25613146	T						3p22.1	rs1052501		41925398	C	1.32 (1.20-1.45) $r^2 = 0.309$	7.47×10^{-9} $D' = 0.643$	1.25 (1.18-1.31) 1.26 (1.20-1.33)	8.19×10^{-17} 8.75×10^{-18}	3	rs6599192	41992408	G	7p15.3	rs4487645	rs4487645	21938240	C	1.38 (1.28-1.50)	3.33×10^{-15}	1.24 (1.19-1.30)	5.30×10^{-25}	3	3q26.2	rs10936599		169492101	C	1.26 (1.18-1.33) $r^2 = 1$	8.70×10^{-14} $D' = 1$	1.20 (1.14-1.25) 1.20 (1.15-1.26)	9.35×10^{-15} 5.94×10^{-15}	4	rs10936600	169514585	A	6p21.3	rs2285803		31107258	T	1.19 (1.13-1.26) $r^2 = 0.94$	9.67×10^{-11} $D' = 0.98$	1.19 (1.14-1.24) 1.20 (1.15-1.25)	1.04×10^{-16} 2.97×10^{-17}	4	rs3132535	31116526	A	17p11.2	rs4273077		16849139	G	1.26 (1.16-1.36) $r^2 = 0.904$	7.67×10^{-9} $D' = 0.958$	1.29 (1.21-1.37) 1.30 (1.23-1.39)	7.35×10^{-15} 3.63×10^{-17}	4	rs34562254	16842991	A	22q13.1	rs877529		39542292	A	1.23 (1.17-1.29) $r^2 = 1$	7.63×10^{-16} $D' = 1$	1.23 (1.19-1.28) 1.23 (1.19-1.28)	6.07×10^{-26} 4.98×10^{-26}	4	rs139402	39546145	C	5q15	rs56219066		95242931	T	1.25 (1.16-1.34) $r^2 = 0.962$	9.6×10^{-10} $D' = 0.981$	1.16 (1.10-1.22) 1.17 (1.12-1.22)	8.06×10^{-8} 1.57×10^{-11}	5	rs1423269	95255724	A	22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$
3p22.1	rs1052501		41925398	C	1.32 (1.20-1.45) $r^2 = 0.309$	7.47×10^{-9} $D' = 0.643$	1.25 (1.18-1.31) 1.26 (1.20-1.33)	8.19×10^{-17} 8.75×10^{-18}	3																																																																																														
		rs6599192	41992408	G						7p15.3	rs4487645	rs4487645	21938240	C	1.38 (1.28-1.50)	3.33×10^{-15}	1.24 (1.19-1.30)	5.30×10^{-25}	3	3q26.2	rs10936599		169492101	C	1.26 (1.18-1.33) $r^2 = 1$	8.70×10^{-14} $D' = 1$	1.20 (1.14-1.25) 1.20 (1.15-1.26)	9.35×10^{-15} 5.94×10^{-15}	4	rs10936600	169514585	A	6p21.3	rs2285803		31107258	T	1.19 (1.13-1.26) $r^2 = 0.94$	9.67×10^{-11} $D' = 0.98$	1.19 (1.14-1.24) 1.20 (1.15-1.25)	1.04×10^{-16} 2.97×10^{-17}	4	rs3132535	31116526	A	17p11.2	rs4273077		16849139	G	1.26 (1.16-1.36) $r^2 = 0.904$	7.67×10^{-9} $D' = 0.958$	1.29 (1.21-1.37) 1.30 (1.23-1.39)	7.35×10^{-15} 3.63×10^{-17}	4	rs34562254	16842991	A	22q13.1	rs877529		39542292	A	1.23 (1.17-1.29) $r^2 = 1$	7.63×10^{-16} $D' = 1$	1.23 (1.19-1.28) 1.23 (1.19-1.28)	6.07×10^{-26} 4.98×10^{-26}	4	rs139402	39546145	C	5q15	rs56219066		95242931	T	1.25 (1.16-1.34) $r^2 = 0.962$	9.6×10^{-10} $D' = 0.981$	1.16 (1.10-1.22) 1.17 (1.12-1.22)	8.06×10^{-8} 1.57×10^{-11}	5	rs1423269	95255724	A	22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$	5.7×10^{-8} $D' = 0.983$	1.11 (1.07-1.16) 1.21 (1.13-1.30)	2.31×10^{-7} 2.58×10^{-8}	5	rs138747	35700488	A						
7p15.3	rs4487645	rs4487645	21938240	C	1.38 (1.28-1.50)	3.33×10^{-15}	1.24 (1.19-1.30)	5.30×10^{-25}	3																																																																																														
3q26.2	rs10936599		169492101	C	1.26 (1.18-1.33) $r^2 = 1$	8.70×10^{-14} $D' = 1$	1.20 (1.14-1.25) 1.20 (1.15-1.26)	9.35×10^{-15} 5.94×10^{-15}	4																																																																																														
		rs10936600	169514585	A						6p21.3	rs2285803		31107258	T	1.19 (1.13-1.26) $r^2 = 0.94$	9.67×10^{-11} $D' = 0.98$	1.19 (1.14-1.24) 1.20 (1.15-1.25)	1.04×10^{-16} 2.97×10^{-17}	4	rs3132535	31116526	A	17p11.2	rs4273077		16849139	G	1.26 (1.16-1.36) $r^2 = 0.904$	7.67×10^{-9} $D' = 0.958$	1.29 (1.21-1.37) 1.30 (1.23-1.39)	7.35×10^{-15} 3.63×10^{-17}	4	rs34562254	16842991	A	22q13.1	rs877529		39542292	A	1.23 (1.17-1.29) $r^2 = 1$	7.63×10^{-16} $D' = 1$	1.23 (1.19-1.28) 1.23 (1.19-1.28)	6.07×10^{-26} 4.98×10^{-26}	4	rs139402	39546145	C	5q15	rs56219066		95242931	T	1.25 (1.16-1.34) $r^2 = 0.962$	9.6×10^{-10} $D' = 0.981$	1.16 (1.10-1.22) 1.17 (1.12-1.22)	8.06×10^{-8} 1.57×10^{-11}	5	rs1423269	95255724	A	22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$	5.7×10^{-8} $D' = 0.983$	1.11 (1.07-1.16) 1.21 (1.13-1.30)	2.31×10^{-7} 2.58×10^{-8}	5	rs138747	35700488	A																													
6p21.3	rs2285803		31107258	T	1.19 (1.13-1.26) $r^2 = 0.94$	9.67×10^{-11} $D' = 0.98$	1.19 (1.14-1.24) 1.20 (1.15-1.25)	1.04×10^{-16} 2.97×10^{-17}	4																																																																																														
		rs3132535	31116526	A						17p11.2	rs4273077		16849139	G	1.26 (1.16-1.36) $r^2 = 0.904$	7.67×10^{-9} $D' = 0.958$	1.29 (1.21-1.37) 1.30 (1.23-1.39)	7.35×10^{-15} 3.63×10^{-17}	4	rs34562254	16842991	A	22q13.1	rs877529		39542292	A	1.23 (1.17-1.29) $r^2 = 1$	7.63×10^{-16} $D' = 1$	1.23 (1.19-1.28) 1.23 (1.19-1.28)	6.07×10^{-26} 4.98×10^{-26}	4	rs139402	39546145	C	5q15	rs56219066		95242931	T	1.25 (1.16-1.34) $r^2 = 0.962$	9.6×10^{-10} $D' = 0.981$	1.16 (1.10-1.22) 1.17 (1.12-1.22)	8.06×10^{-8} 1.57×10^{-11}	5	rs1423269	95255724	A	22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$	5.7×10^{-8} $D' = 0.983$	1.11 (1.07-1.16) 1.21 (1.13-1.30)	2.31×10^{-7} 2.58×10^{-8}	5	rs138747	35700488	A																																										
17p11.2	rs4273077		16849139	G	1.26 (1.16-1.36) $r^2 = 0.904$	7.67×10^{-9} $D' = 0.958$	1.29 (1.21-1.37) 1.30 (1.23-1.39)	7.35×10^{-15} 3.63×10^{-17}	4																																																																																														
		rs34562254	16842991	A						22q13.1	rs877529		39542292	A	1.23 (1.17-1.29) $r^2 = 1$	7.63×10^{-16} $D' = 1$	1.23 (1.19-1.28) 1.23 (1.19-1.28)	6.07×10^{-26} 4.98×10^{-26}	4	rs139402	39546145	C	5q15	rs56219066		95242931	T	1.25 (1.16-1.34) $r^2 = 0.962$	9.6×10^{-10} $D' = 0.981$	1.16 (1.10-1.22) 1.17 (1.12-1.22)	8.06×10^{-8} 1.57×10^{-11}	5	rs1423269	95255724	A	22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$	5.7×10^{-8} $D' = 0.983$	1.11 (1.07-1.16) 1.21 (1.13-1.30)	2.31×10^{-7} 2.58×10^{-8}	5	rs138747	35700488	A																																																							
22q13.1	rs877529		39542292	A	1.23 (1.17-1.29) $r^2 = 1$	7.63×10^{-16} $D' = 1$	1.23 (1.19-1.28) 1.23 (1.19-1.28)	6.07×10^{-26} 4.98×10^{-26}	4																																																																																														
		rs139402	39546145	C						5q15	rs56219066		95242931	T	1.25 (1.16-1.34) $r^2 = 0.962$	9.6×10^{-10} $D' = 0.981$	1.16 (1.10-1.22) 1.17 (1.12-1.22)	8.06×10^{-8} 1.57×10^{-11}	5	rs1423269	95255724	A	22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$	5.7×10^{-8} $D' = 0.983$	1.11 (1.07-1.16) 1.21 (1.13-1.30)	2.31×10^{-7} 2.58×10^{-8}	5	rs138747	35700488	A																																																																				
5q15	rs56219066		95242931	T	1.25 (1.16-1.34) $r^2 = 0.962$	9.6×10^{-10} $D' = 0.981$	1.16 (1.10-1.22) 1.17 (1.12-1.22)	8.06×10^{-8} 1.57×10^{-11}	5																																																																																														
		rs1423269	95255724	A						22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$	5.7×10^{-8} $D' = 0.983$	1.11 (1.07-1.16) 1.21 (1.13-1.30)	2.31×10^{-7} 2.58×10^{-8}	5	rs138747	35700488	A																																																																																	
22q13	rs138740		35699582	C	1.18 (1.11-1.25) $r^2 = 0.955$	5.7×10^{-8} $D' = 0.983$	1.11 (1.07-1.16) 1.21 (1.13-1.30)	2.31×10^{-7} 2.58×10^{-8}	5																																																																																														
		rs138747	35700488	A																																																																																																			

Supplementary Table 3: Strongest association signals from previously published risk loci discovered in European populations³⁻⁵. Shown for each region are the published SNP, the most associated variant within a 500kb window in the imputation and the odds ratio and *P* values associated with each, along with the linkage disequilibrium metrics between the SNPs. Meta-analysis for current GWAS was undertaken using the inverse-variance approach under a fixed effects model.

SNP		rs34229995	rs9372120	rs7781265	rs1948915	rs2811710	rs2790457	rs7193541	rs6066835
Locus		6p22.3	6q21	7q36.1	8q24.21	9p21.3	10p12.1	16q23.1	20q13.13
Position (bp, hg19)		15,244,018	106,667,535	150,950,940	128,222,421	21,991,923	28,856,819	74,664,743	47,355,009
Allele A		C	T	C	T	A	A	C	T
Allele B (Risk Allele)		G	G	T	C	G	G	T	C
UK	Cases RAF	0.024	0.236	0.128	0.332	0.666	0.751	0.628	0.100
	Controls RAF	0.019	0.211	0.117	0.319	0.634	0.733	0.584	0.083
	OR	1.28	1.16	1.11	1.06	1.16	1.10	1.20	1.25
	<i>P</i> value	0.0525	6.72×10^{-4}	0.0578	0.109	1.24×10^{-4}	0.0149	5.97×10^{-7}	3.83×10^{-4}
Sweden/ Norway	Cases RAF	0.045	0.243		0.386	0.681	0.750	0.598	0.097
	Controls RAF	0.033	0.217		0.363	0.660	0.735	0.572	0.075
	OR	1.44	1.22		1.12	1.14	1.11	1.07	1.38
	<i>P</i> value	4.37×10^{-4}	9.96×10^{-6}	Info<0.8	2.97×10^{-3}	1.38×10^{-3}	0.0160	0.0612	6.20×10^{-6}
Germany	Cases RAF	0.039	0.226	0.144	0.357	0.678	0.757	0.596	0.095
	Controls RAF	0.028	0.207	0.118	0.331	0.654	0.728	0.590	0.077
	OR	1.49	1.18	1.33	1.22	1.15	1.15	1.02	1.24
	<i>P</i> value	9.03×10^{-3}	0.0124	3.22×10^{-4}	3.13×10^{-4}	0.0119	0.0224	0.645	0.0240
Iceland	Cases RAF								
	Controls RAF								
	OR	1.32	1.19	1.14	1.21	1.06	1.19	1.14	1.08
	<i>P</i> value	0.0110	0.0501	0.190	0.0153	0.440	0.0402	0.0907	0.535
USA	Cases RAF	0.036	0.239	0.159	0.362	0.669	0.761	0.604	0.105
	Controls RAF	0.023	0.202	0.120	0.313	0.637	0.732	0.584	0.082
	OR	1.61	1.25	1.45	1.25	1.16	1.17	1.09	1.33
	<i>P</i> value	0.0110	2.46×10^{-3}	6.56×10^{-5}	5.35×10^{-4}	0.0258	0.0232	0.166	7.11×10^{-3}
Netherlands	Cases RAF	0.031	0.239	0.137	0.363	0.682	0.749	0.616	0.081
	Controls RAF	0.024	0.204	0.124	0.329	0.658	0.742	0.570	0.085
	OR	1.32	1.23	1.12	1.17	1.12	1.04	1.21	0.95
	<i>P</i> value	0.192	9.45×10^{-3}	0.240	0.0258	0.113	0.620	4.36×10^{-3}	0.687
Meta	OR	1.40	1.20	1.20	1.14	1.14	1.12	1.12	1.24
	(95% CI)	(1.25-1.58)	(1.14-1.26)	(1.12-1.29)	(1.10-1.19)	(1.09-1.19)	(1.07-1.17)	(1.08-1.17)	(1.16-1.33)
	<i>P</i> value	1.76×10^{-8}	8.72×10^{-14}	1.82×10^{-7}	3.14×10^{-10}	6.50×10^{-10}	8.44×10^{-7}	1.14×10^{-8}	1.16×10^{-9}

Supplementary Table 4: Summary statistics for novel variants showing an association with multiple myeloma risk in the GWAS meta-analysis at $P < 1.0 \times 10^{-6}$. Showing SNPs which were genome-wide significant after replication. Odds ratios derived with respect to the risk allele. Cases RAF, risk allele frequency in discovery cases. Controls RAF, risk allele frequency in discovery controls. Shown are discovery association P values for individual studies (logistic regression) and from meta-analysis of the six GWAS datasets (inverse-variance approach under a fixed effects model).

SNP		rs78311596	rs72665486	rs1034447	rs17507636	rs28571765	rs76601148	rs8058578	rs6071887
Locus		1p13.2	4q22.1	6q27	7q22.3	8q11.23	13q13.3	16p11.2	20q12
Position (bp, hg19)		113,300,945	91,913,730	164,613,457	106,291,118	55,449,281	36,746,056	30,726,248	38,621,456
Allele A		A	T	C	T	C	A	C	G
Allele B (Risk Allele)		T	G	T	C	T	G	T	A
UK	Cases RAF	0.029	0.882	0.163	0.760	0.808	0.049	0.280	0.258
	Controls RAF	0.022	0.874	0.148	0.745	0.794	0.041	0.255	0.238
	OR	1.34	1.07	1.13	1.08	1.09	1.07	1.14	1.11
	<i>P</i> value	0.0138	0.180	0.0148	0.0476	0.0532	0.0152	1.34×10^{-3}	8.42×10^{-3}
Sweden/ Norway	Cases RAF		0.907	0.158	0.778	0.797	0.051	0.283	0.225
	Controls RAF		0.886	0.127	0.765	0.773	0.038	0.261	0.208
	OR		1.25	1.27	1.09	1.13	1.39	1.11	1.11
	<i>P</i> value	MAF<0.005	2.14×10^{-4}	1.40×10^{-5}	0.0436	8.80×10^{-3}	6.57×10^{-4}	0.0126	0.0199
Germany	Cases RAF	0.020	0.907	0.142	0.745	0.825	0.039	0.281	0.220
	Controls RAF	0.016	0.883	0.138	0.714	0.812	0.033	0.261	0.213
	OR	1.449	1.36112	1.0950	1.13882	1.09694	1.4424	1.0979	1.06395
	<i>P</i> value	0.0733	2.9×10^{-4}	0.238	0.0281	0.175	0.0111	0.115	0.331
Iceland	Cases RAF								
	Controls RAF								
	OR	1.71	1.21	1.13	1.06	1.30	1.07	1.19	1.12
	<i>P</i> value	1.47×10^{-3}	0.0970	0.226	0.491	3.75×10^{-3}	0.692	0.0350	0.213
USA	Cases RAF	0.027	0.887	0.144	0.785	0.826	0.051	0.273	0.267
	Controls RAF	0.017	0.884	0.136	0.737	0.789	0.041	0.254	0.231
	OR	1.73	1.02	1.06	1.30	1.26	1.28	1.10	1.22
	<i>P</i> value	0.0132	0.811	0.489	2.03×10^{-4}	2.31×10^{-3}	0.0963	0.160	4.90×10^{-3}
Netherlands	Cases RAF	0.026	0.896	0.153	0.764	0.825	0.059	0.259	0.251
	Controls RAF	0.021	0.878	0.141	0.735	0.803	0.043	0.245	0.221
	OR	1.26	1.19	1.10	1.16	1.16	1.45	1.07	1.17
	<i>P</i> value	0.321	0.0870	0.300	0.0440	0.0776	0.0206	0.259	0.0380
Meta	OR	1.46	1.17	1.15	1.12	1.14	1.31	1.12	1.12
	(95% CI)	(1.25-1.69)	(1.10-1.24)	(1.09-1.22)	(1.07-1.17)	(1.08-1.19)	(1.19-1.45)	(1.07-1.17)	(1.07-1.18)
	<i>P</i> value	9.40×10^{-7}	4.81×10^{-7}	8.21×10^{-7}	5.54×10^{-7}	1.30×10^{-7}	5.90×10^{-8}	5.57×10^{-7}	7.08×10^{-7}

Supplementary Table 5: Summary statistics for novel variants showing an association with multiple myeloma risk in the GWAS meta-analysis at $P < 1.0 \times 10^{-6}$. Showing SNPs which were not genome-wide significant after replication. Odds ratios derived with respect to the risk allele. Cases RAF, risk allele frequency in discovery cases. Controls RAF, risk allele frequency in discovery controls. Shown are discovery association P values for individual studies (logistic regression) and from meta-analysis of the six GWAS datasets (inverse-variance approach under a fixed effects model).

SNP		rs34229995	rs9372120	rs7781265	rs1948915	rs2811710	rs2790457	rs7193541	rs6066835
UK Cases	AA	164/164	105/105	126/127	77/77	61/62	97/97	Genotyped	130/130
	Aa	7/7	57/59	39/40	68/68	71/73	65/65		41/41
	aa	2/2	7/7	3/3	13/14	23/23	12/12		2/2
	r^2	1.00	1.0	0.97	0.99	0.98	1.00		1.00
UK Controls	AA	146/146	110/110	123/123	75/77	69/70	85/87		142/142
	Aa	16/16	52/52	30/32	64/64	62/63	63/64		21/21
	aa	1/1	1/1	2/2	12/13	15/16	12/12		NA
	r^2	1.00	1.00	0.97	0.98	0.98	0.99		1.00
Germany Cases	AA	314/314	193/193	250/254	135/136	146/153	184/185		279/279
	Aa	20/20	124/125	68/70	142/148	134/138	124/124		53/55
	aa	1/1	12/12	6/6	36/38	26/27	26/26		3/3
	r^2	1.00	0.99	0.95	0.97	0.94	1.00		0.98
USA Cases	AA	692/692	428/431	494/506	307/308	308/314	420/421		591/591
	Aa	50/50	268/269	149/159	320/322	296/299	276/278		143/143
	aa	2/2	41/41	13/14	103/104	72/74	38/38		6/6
	r^2	1.00	0.98	0.86	0.99	0.96	0.99		1.0

Supplementary Table 6: Concordance between directly sequenced and imputed genotype. Showing SNPs which were genome-wide significant after replication. AA, major homozygote; AB, heterozygote; BB, minor homozygote. r^2 indicates Pearson product-moment correlation coefficient between imputed and sequenced genotype.

Study	Genotype	rs78311596	rs72665486	rs1034447	rs17507636	rs28571765	rs76601148	rs8058578	rs6071887
UK Cases	AA	166/166	127/127	120/120	Genotyped	108/108	151/151	Genotyped	97/97
	Aa	8/9	35/35	48/48		61/62	22/22		59/59
	aa	N/A	3/3	6/6		4/4	2/2		15/15
	r^2	0.94	1.00	1.00		0.99	1.00		1.00
UK Controls	AA	142/142	124/125	126/126	101/101	105/105			85/85
	Aa	17/18	37/37	35/35	52/52	56/56			70/70
	aa	2/2	1/1	1/1	8/8	2/2			3/3
	r^2	0.98	1.00	1.00	1.00	1.00			1.00
Germany Cases	AA	318/318	138/138	246/247		228/229	317/318		189/202
	Aa	9/9	32/32	87/87		91/91	19/20		109/109
	aa	NA	NA	5/5		9/9	NA		14/14
	r^2	1.00	1.00	0.99		0.99	0.97		0.91
USA Cases	AA	704/705	587/587	539/540		512/512	672/672		396/396
	Aa	28/34	154/154	189/191		193/193	71/71		93/293
	aa	1/1	8/9	10/10		29/29	2/2		49/49
	r^2	0.80	0.99	0.98		1.0	1.0		1.0

Supplementary Table 7: Concordance between directly sequenced and imputed genotype. Showing SNPs which were not genome-wide significant after replication. AA, major homozygote; AB, heterozygote; BB, minor homozygote. r^2 indicates Pearson product-moment correlation coefficient between imputed and sequenced genotype.

SNP		rs34229995	rs9372120	rs7781265	rs1948915	rs2811710	rs2790457	rs7193541	rs6066835
Locus		6p22.3	6q21	7q36.1	8q24.21	9p21.3	10p12.1	16q23.1	20q13.13
Position (bp, hg19)		15,244,018	106,667,535	150,950,940	128,222,421	21,991,923	28,856,819	74,664,743	47,355,009
Allele A		C	T	C	T	A	A	C	T
Allele B (Risk Allele)		G	G	T	C	G	G	T	C
UK	Cases RAF	0.022	0.234	0.136	0.332	0.666	0.752	0.63	0.101
	Controls RAF	0.020	0.210	0.113	0.309	0.634	0.733	0.58	0.083
	OR	1.14	1.15	1.23	1.11	1.12	1.07	1.11	1.37
	<i>P</i> value	0.576	0.091	0.043	0.151	0.106	0.423	0.147	0.008
Sweden/Norway	Cases RAF	0.010	0.210	0.143	0.404	0.710	0.758	0.618	0.070
	Controls RAF	0.020	0.201	0.125	0.366	0.666	0.729	0.583	0.071
	OR	0.59	1.052	1.171	1.17	1.23	1.17	1.156	0.99
	<i>P</i> value	0.128	0.677	0.267	0.106	0.053	0.164	0.142	0.974
Germany	Cases RAF	0.024	0.211	0.134	0.350	0.669	0.759	0.614	0.102
	Controls RAF	0.015	0.197	0.127	0.330	0.630	0.727	0.565	0.805
	OR	1.60	1.09	1.06	1.10	1.19	1.190	1.22	1.29
	<i>P</i> value	0.03	0.26	0.51	0.16	0.009	0.02	0.001	0.015
Denmark	Cases RAF	0.012	0.229	0.147	0.365	0.698	0.751	0.616	0.121
	Controls RAF	0.012	0.199	0.121	0.366	0.651	0.733	0.590	0.074
	OR	1.062	1.19	1.25	1.00	1.24	1.10	1.114	1.71
	<i>P</i> value	0.870	0.081	0.069	0.960	0.019	0.349	0.228	1×10^{-4}
Discovery phase + replication	OR	1.37	1.18	1.19	1.13	1.15	1.12	1.13	1.26
	<i>P</i> value	1.31×10^{-8}	9.09×10^{-15}	9.71×10^{-9}	4.20×10^{-11}	1.72×10^{-13}	1.77×10^{-8}	5.00×10^{-12}	1.36×10^{-13}

Supplementary Table 8: Replication of top association signals. Showing SNPs which were genome-wide significant after replication. Cases RAF, risk allele frequency of replication cases; Control RAF, risk allele frequency of replication controls. *P* values are shown for each replication series (logistic regression) and for the meta-analysis of the 6 discovery datasets and the four replication series combined (inverse-variance approach under a fixed effects model).

SNP		rs72665486	rs1034447	rs17507636	rs28571765	rs76601148	rs8058578	rs6071887
Locus		4q22.1	6q27	7q22.3	8q11.23	13q13.3	16p11.2	20q12
Position (bp, hg19)		91,913,730	164,613,457	106,291,118	55,449,281	36,746,056	30,726,248	38,621,456
Allele A		T	C	T	C	A	C	G
Allele B (Risk Allele)		G	T	C	T	G	T	A
UK	Cases RAF	0.882	0.163	0.760	0.808	0.049	0.280	0.258
	Controls RAF	0.874	0.148	0.745	0.794	0.041	0.255	0.238
	OR	1.07	1.02	1.09	1.15	1.07	1.15	1.10
	<i>P</i> value	0.440	0.812	0.312	0.098	0.694	0.064	0.214
Sweden/Norway	Cases RAF	0.871	0.155	0.784	0.790	0.040	Genotyping	0.188
	Controls RAF	0.879	0.139	0.764	0.785	0.039	failed	0.212
	OR	0.93	1.14	1.27	1.03	1.04		0.86
	<i>P</i> value	0.639	0.400	0.308	0.784	0.882		0.236
Germany	Cases RAF	0.893	0.138	0.760	0.810	0.366	0.296	0.232
	Controls RAF	0.884	0.131	0.735	0.807	0.410	0.304	0.216
	OR	1.10	1.06	1.15	1.02	0.89	0.96	1.10
	<i>P</i> value	0.34	0.51	0.06	0.81	0.47	0.59	0.23
Denmark	Cases RAF	0.884	0.168	0.734	0.766	0.033	Genotyping	0.223
	Controls RAF	0.889	0.146	0.751	0.80	0.039	failed	0.215
	OR	0.95	1.18	0.93	0.79	0.85		1.05
	<i>P</i> value	0.709	0.171	0.434	0.0182	0.455		0.654
Discovery phase + replication	OR	1.14	1.13	1.11	1.10	1.23	1.10	1.11
	<i>P</i> value	2.15×10^{-6}	4.81×10^{-7}	1.58×10^{-7}	1.58×10^{-6}	4.46×10^{-6}	9.68×10^{-7}	6.47×10^{-7}

Supplementary Table 9: Replication of top association signals. Showing SNPs which were not genome-wide significant after replication. Cases RAF, risk allele frequency of replication cases; Control RAF, risk allele frequency of replication controls. *P* values are shown for each replication series (logistic regression) and for the meta-analysis of the 6 discovery datasets and the four replication series combined (inverse-variance approach under a fixed effects model).

Cancer	SNP	Reference	Previously Reported		Current MM GWAS		LD with rs1948915	
			Risk Allele	<i>P</i> value	Risk Allele	<i>P</i> value	<i>r</i> ²	<i>D'</i>
Hodgkin's Lymphoma	rs2019960	6	C	7×10^{-8}	C	0.024	1.9×10^{-5}	-0.012
Colorectal Cancer	rs6983267	7	G	5×10^{-14}	T	0.17	8.7×10^{-4}	-0.046
Prostate Cancer	rs1447295	8	A	6×10^{-18}	C	0.55	1.6×10^{-4}	0.055
Prostate Cancer	rs12682344	8	G	5×10^{-12}	G	0.48	5.2×10^{-3}	-0.57
Prostate Cancer (early onset)	rs10505477	9	A	9×10^{-9}	G	0.11	1.1×10^{-3}	-0.051
Colorectal Cancer	rs10505477	10	T	8×10^{-13}	G	0.11	1.1×10^{-3}	-0.051
Breast Cancer (early onset)	rs2392780	11		1×10^{-8}	G	0.86	1.0×10^{-3}	0.039
Renal cell carcinoma	rs6470589	12	G	5×10^{-11}	G	0.82		
Bladder Cancer	rs9642880	13	T	4×10^{-38}	T	0.17	3.9×10^{-5}	8.6×10^{-3}
Chronic lymphocytic leukemia	rs2466035	14	C	2×10^{-8}	C	3.6×10^{-3}	0.15	0.39
Breast Cancer	rs13281615	15	G	1×10^{-17}	G	0.076	6.8×10^{-4}	-0.046
Breast Cancer	rs11780156	15	T	3×10^{-11}	T	0.22	2.8×10^{-5}	0.0076
Ovarian Cancer	rs10088218	16		1×10^{-17}	A	0.86	1.0×10^{-3}	-0.13
Prostate Cancer	rs6983561	17	C	4×10^{-13}	C	0.47	4.9×10^{-3}	-0.55
Prostate Cancer	rs13254738	18	C	4×10^{-10}	A	0.43	0.030	-0.17
Prostate Cancer	rs4242384	19	C	3×10^{-16}	A	0.57	2.1×10^{-4}	0.063
Prostate Cancer	rs1016343	19	T	4×10^{-10}	T	0.63	0.015	0.16
Glioma	rs4295627	20	G	5×10^{-21}	G	5.4×10^{-3}	6.3×10^{-4}	0.0047
Breast Cancer	rs1562430	21	A	3×10^{-11}	C	0.65	8.9×10^{-4}	0.037
Prostate Cancer	rs1456315	22	A	2×10^{-29}	C	0.26	0.028	-0.17
Prostate Cancer	rs7837688	22	T	1×10^{-25}	G	0.68	3.7×10^{-4}	0.083
Prostate Cancer	rs16902094	23	G	6×10^{-15}	G	0.74	1.9×10^{-4}	-0.049
Prostate Cancer	rs16901979	23	A	3×10^{-14}	A	0.50	5.2×10^{-3}	-0.57
Prostate Cancer	rs445114	23	T	5×10^{-10}	C	0.13	1.6×10^{-3}	-0.076
Glioma	rs891835	24	G	8×10^{-11}	G	0.039	2.1×10^{-5}	5.8×10^{-3}
Colorectal Cancer	rs7014346	25	A	9×10^{-26}	G	7.9×10^{-4}	2.2×10^{-4}	0.028
Prostate Cancer	rs6983267	26	G	7×10^{-12}	T	0.17	8.7×10^{-4}	-0.046
Glioma	rs55705857	27	A	2.24×10^{-38}	A	9.5×10^{-4}	1.2×10^{-5}	-0.017

Supplementary Table 10: Previously reported 8q24.21 cancer associated SNPs and their LD with rs1948915⁶⁻²⁷.

Cancer	SNP	Reference	Previously Reported		Current MM GWAS		LD with rs2811710	
			Risk Allele	<i>P</i> value	Risk Allele	<i>P</i> value	<i>r</i> ²	<i>D</i> '
Basal cell carcinoma	rs2151280	28	G	3×10^{-10}	A	0.0127	4.14×10^{-4}	-0.0280
ALL	rs3731249	29	T	9.4×10^{-23}	T	0.957	0.0385	0.920
CLL	rs1679013	30	C	1×10^{-8}	C	6.54×10^{-3}	4.27×10^{-3}	0.0791
Glioma	rs4977756	31	G	1×10^{-8}	A	0.368	0.0513	0.360
Melanoma	rs7023329	32	A	7×10^{-9}	G	0.0822	1.06×10^{-4}	-0.0131
Glioma	rs2157719	20	C	5×10^{-16}	T	0.399	0.0485	0.328
Glioma (High Grade)	rs1412829	33	C	2×10^{-10}	C	0.285	0.0434	-0.317

Supplementary Table 11: Previously reported 9p21.3 cancer associated SNPs and their LD with rs2811710^{20,28-33}.

		rs34229995	rs9372120	rs7781265	rs1948915	rs2811710	rs2790457	rs7193541	rs6066835
UK	Beta	0.44	0.64	-0.093	-0.43	-0.20	-0.36	-0.022	-0.075
	<i>P</i> value	0.67	0.076	0.84	0.19	0.56	0.31	0.94	0.88
Sweden/ Norway	Beta	1.88	1.31	NA	0.33	-0.054	0.60	-0.30	1.82
	<i>P</i> value	0.070	9.5×10^{-3}	NA	0.47	0.91	0.23	0.51	0.01
Germany	Beta	0.068	0.15	-0.41	0.34	0.40	0.24	0.22	0.14
	<i>P</i> value	0.94	0.73	0.43	0.35	0.32	0.57	0.56	0.82
USA	Beta	0.93	-1.18	0.023	0.28	0.017	-0.51	0.55	-0.71
	<i>P</i> value	0.45	0.032	0.97	0.56	0.97	0.36	0.25	0.35
Netherlands	Beta	-0.96	-0.11	-0.12	-0.046	-0.45	-0.33	-0.47	-0.21
	<i>P</i> value	0.16	0.67	0.95	0.84	0.049	0.16	0.025	0.57

Supplementary Table 12: Relationship between SNP genotype and age at diagnosis. Analysis based on Beta values calculated from linear regression on the discovery phase data sets from UK (cases=2282, controls=5197), Sweden/Norway (cases=1714, controls=10391), Germany (cases=1508, controls=2107), USA (cases=780, controls=1857) and Netherlands (cases=555, controls=2669).

		rs34229995	rs9372120	rs7781265	rs1948915	rs2811710	rs2790457	rs7193541	rs6066835
UK	Beta	-0.052	-0.013	-0.11	0.036	-0.028	0.12	0.021	0.069
	<i>P</i> value	0.79	0.85	0.22	0.58	0.67	0.081	0.74	0.49
Sweden/ Norway	Beta	0.26	-0.020	NA	-0.083	-0.042	-0.12	-0.11	0.11
	<i>P</i> value	0.12	0.81	NA	0.25	0.58	0.12	0.13	0.34
Germany	Beta	-0.078	-0.11	0.021	0.049	-0.028	-0.06	-0.012	-0.24
	<i>P</i> value	0.69	0.22	0.84	0.53	0.74	0.45	0.88	0.06
USA	Beta	-0.43	0.16	-0.061	0.019	-0.13	-0.040	-0.046	-0.60
	<i>P</i> value	0.15	0.20	0.69	0.86	0.24	0.75	0.67	1.7×10^{-3}
Netherlands	Beta	-0.11	-0.13	0.077	-0.11	0.017	-0.012	0.028	-0.068
	<i>P</i> value	0.95	0.88	0.46	0.99	0.86	0.90	0.78	0.55

Supplementary Table 13: Relationship between SNP genotype and sex of multiple myeloma cases. Analysis based on Beta values calculated from logistic regression on the discovery phase data sets from UK (cases=2282, controls=5197), Sweden/Norway (cases=1714, controls=10391), Germany (cases=1508, controls=2107), USA (cases=780, controls=1857) and Netherlands (cases=555, controls=2669).

			rs34229995	rs9372120	rs7781265	rs1948915	rs2811710	rs2790457	rs7193541	rs6066835
Hyperdiploid	UK (702)	Beta	0.0285	-0.111	-6.14×10^{-3}	-0.0441	0.0191	-0.142	-0.0374	0.369
		P value	0.892	0.140	0.950	0.522	0.783	0.0555	0.569	5.74×10^{-4}
	Germany (661)	Beta	-0.152	-0.0373	7.33×10^{-3}	0.0802	-1.37×10^{-3}	-0.0318	-0.0804	0.0649
		P value	0.435	0.677	0.944	0.303	0.987	0.713	0.281	0.600
	Meta	Beta	-0.0686	-0.0805	1.17×10^{-4}	0.0104	0.0105	-0.0953	-0.0562	0.239
		P value	0.630	0.162	0.999	0.840	0.843	0.0901	0.255	3.20×10^{-3}
Translocation	UK (571)	Beta	0.0181	0.155	-0.230	0.0758	0.116	-0.137	0.0190	-0.0618
		P value	0.936	0.0517	0.0272	0.302	0.118	0.0818	0.786	0.588
	Germany (768)	Beta	0.187	-8×10^{-4}	-0.0232	-0.0127	-6.96×10^{-3}	-0.0771	-0.0657	-0.151
		P value	0.331	0.992	0.824	0.870	0.931	0.370	0.375	0.219
	Meta	Beta	0.116	0.0857	-0.127	0.0338	0.0596	-0.110	-0.0210	-0.103
		P value	0.429	0.150	0.0854	0.526	0.274	0.0589	0.680	0.217
t(11;14)	UK (231)	Beta	0.0335	0.186	-0.380	0.0845	0.119	-0.322	0.0191	0.171
		P value	0.918	0.104	0.0110	0.423	0.261	4.37×10^{-3}	0.850	0.296
	Germany (277)	Beta	-0.232	0.0900	-0.0818	-0.195	-0.0151	0.0881	-0.0224	-0.130
		P value	0.350	0.434	0.543	0.0510	0.885	0.427	0.815	0.409
	Meta	Beta	-0.134	0.138	-0.215	-0.0628	0.0509	-0.113	-2.69×10^{-3}	0.0151
		P value	0.497	0.0886	0.0311	0.387	0.494	0.153	0.969	0.894
t(4;14)	UK (170)	Beta	-0.392	0.0396	-0.0224	0.154	0.0172	0.176	0.0733	-0.351
		P value	0.292	0.764	0.897	0.204	0.887	0.177	0.526	0.0622
	Germany (142)	Beta	0.521	-0.110	0.206	0.0281	-0.170	-0.122	-0.135	-0.129
		P value	0.112	0.472	0.245	0.831	0.219	0.407	0.289	0.541
	Meta	Beta	0.121	-0.0243	0.0887	0.0963	-0.0644	0.0451	-0.0210	-0.252
		P value	0.622	0.808	0.474	0.281	0.480	0.644	0.806	0.0724
t(14;16)	UK (24)	Beta	1.67	0.309	-0.210	-0.139	-0.189	-0.342	0.0767	-0.896
		P value	0.0778	0.361	0.635	0.657	0.541	0.305	0.798	0.0637
	Germany (29)	Beta	0.384	-0.182	0.205	0.0490	0.235	-0.283	0.0260	-0.487
		P value	0.586	0.587	0.595	0.863	0.420	0.367	0.923	0.284
	Meta	Beta	0.843	0.0659	0.0263	-0.0362	-0.0356	0.311	-0.0486	-0.680
		P value	0.136	0.784	0.928	0.863	0.867	0.174	0.809	0.0403

Supplementary Table 14: Relationship between SNP genotype and multiple myeloma subtype. Case-only analysis; Beta values obtained from logistic regression. FISH and ploidy classification of UK and German samples were determined as previously described³⁴⁻³⁶. Number of samples in each cohort indicated in brackets.

SNP		rs34229995	rs9372120	rs7781265	rs1948915	rs2811710	rs2790457	rs7193541	rs6066835
Locus		6p22.3	6q21	7q36.1	8q24.21	9p21.3	10p12.1	16q23.1	20q13.13
Position (bp, hg19)		15,244,018	106,667,535	150,950,940	128,222,421	21,991,923	28,856,819	74,664,743	47,355,009
Allele A		C	T	C	T	A	A	C	T
Allele B (Risk Allele)		G	G	T	C	G	G	T	C
Nearest Gene		<i>JARID2</i>	<i>ATG5</i>	<i>SMARCD3</i>	<i>CCAT1</i>	<i>CDKN2A</i>	<i>WAC</i>	<i>RFWD3</i>	<i>PREX1</i>
UK-MyIX	HR	1.48	0.82	0.83	0.98	1.07	0.91	0.94	0.97
	(95%CI)	(1.03-2.12)	(0.71-0.95)	(0.68-1.01)	(0.86-1.13)	(0.94-1.22)	(0.79-1.05)	(0.83-1.07)	(0.79-1.19)
	<i>P</i> value	0.035	0.008	0.06	0.827	0.31	0.208	0.344	0.775
UK-MyXI	HR	0.81	1.19	1.12	1.05	0.91	0.92	1.05	0.88
	(95%CI)	(0.40-1.64)	(0.96-1.47)	(0.84-1.49)	(0.85-1.29)	(0.73-1.13)	(0.73-1.16)	(0.87-1.29)	(0.63-1.25)
	<i>P</i> value	0.555	0.117	0.423	0.679	0.409	0.484	0.61	0.484
GER-GMMG	HR	0.89	1.02	1.19	1.04	0.77	0.88	1.01	1.09
	(95%CI)	(0.49-1.63)	(0.76-1.35)	(0.89-1.60)	(0.83-1.31)	(0.59-1.01)	(0.67-1.17)	(0.78-1.28)	(0.77-1.54)
	<i>P</i> value	0.710	0.918	0.243	0.731	0.06	0.395	0.943	0.634
US-UAMS	HR	1.01	0.96	1.16	0.96	1.07	0.94	1.06	1.11
	(95%CI)	(0.68-1.51)	(0.80-1.5)	(0.93-1.45)	(0.82-1.12)	(0.87-1.22)	(0.78-1.14)	(0.91-1.24)	(0.87-1.42)
	<i>P</i> value	0.958	0.631	0.174	0.591	0.650	0.552	0.430	0.410
Combined	HR	1.13 (0.90-	0.94 (0.86-	1.02 (0.91-	0.99 (0.91-	1.01 (0.92-	0.92 (0.83-	1.00 (0.92-	1.01 (0.89-
	(95%CI)	1.43)	1.04)	1.15)	1.08)	1.10)	1.01)	1.09)	1.15)
	<i>I</i> ²	22	64	55	0	50	0	50	0
	<i>P</i> value	0.295	0.206	0.698	0.894	0.85	0.084	0.845	0.866

Supplementary Table 15: Relationship between genome-wide significant SNPs genotype and patient overall survival³⁷. Data from: 1,165 cases from the UK MRC Myeloma-IX trial (UK-MyIX); 877 MM cases from the UK MRC Myeloma-XI trial (UK-MyXI); 511 of the patients recruited to the German-GWAS (GER-GMMG); 703 MM cases in the UAMS Myeloma Institute for Research and Therapy GWAS (US-UAMS). *P* values calculated from Cox regression analysis.

SNP	Gene	Non silent mutations	Silent mutations	Noncoding mutation	<i>P</i> value	<i>q</i> value (FDR)
rs34229995	<i>DTNBP1</i>	0	1	0	0.184	1
rs34229995	<i>JARID2</i>	0	2	0	0.910	1
rs9372120	<i>ATG5</i>	0	1	0	1	1
rs7781265	<i>SMARCD3</i>	0	0	0	1	1
rs7781265	<i>CHPF2</i>	0	1	0	0.983	1
rs2811710	<i>CDKN2A</i>	0	0	0	1	1
rs2790457	<i>WAC</i>	0	1	0	0.760	1
rs7193541	<i>RFWD3</i>	0	0	0	1	1
rs6066835	<i>PREX1</i>	0	0	0	1	1

Supplementary Table 16: Mutation significance analysed with MutSigCV³⁸ on 463 MM exomes. The frequency of somatic mutation in genes annotated by GWAS signals was derived from tumor whole exome sequencing of Myeloma XI trial patients.

SNP	Locus	Gene 1	Gene 2	Pearson's Correlation (95% CI)	P value
rs9372120	6q21	<i>ATG5</i>	<i>PRDM1</i>	0.50 (0.29-0.66)	1.99×10^{-5}
rs2811710	9p21.3	<i>MTAP</i>	<i>CDKN2A</i>	0.54 (0.34-0.69)	3.13×10^{-6}
rs2811710	9p21.3	<i>CDKN2A</i>	<i>CDKN2B</i>	0.53 (0.33-0.68)	4.99×10^{-6}

Supplementary Table 17: A pair-wise correlation between gene expression (RNA-seq) within TADs. Analysis based on RNAseq transcript counts from 66 MM cell lines from the Keat's lab Data Repository³⁹ (Pearson product-moment correlation coefficient).

SNP	Reaction	Conditions		Sequence
rs34229995	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTGCCCATGGATCCTCCC
			Kaspar A2	GAAGGTGGAGTCAACGGATTCTGCCCATGGATCCTCCG
			Kaspar C	CACTTCACAATTGGAAAAATAAATGATCTT
	Sanger seq	Std	Seq F	ATTCCCAGTATGCCAGTCTT
			Seq R	AGGCACGTATTGGAAGCTT
rs9372120	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTCCTCCCTGGGTTACCAAATTCTC
			Kaspar A2	GAAGGTGGAGTCAACGGATTACCTCCTGGGTTACCAAATTCTA
			Kaspar C	ACCATTTTATATCAGACTCTCAAACATCAG
	Sanger seq	Std	Seq F	TGCACATTAACTCAGCCACA
			Seq R	GGAAACCGTATTTGAGTAGACCA
rs7781265	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTCCAATTCAGGGTCCCTGGAT
			Kaspar A2	GAAGGTGGAGTCAACGGATTCCAATTCAGGGTCCCTGGAC
			Kaspar C	CCCACCTTCCAGGGTGTGGCT
	Sanger seq	Std	Seq F	CCCATTAATGAGGTGGTCGC
			Seq R	CGATCCCCTTGTGTTTGCC
rs1948915	Kaspar	Std42plus5	Kaspar A1	GAAGGTGACCAAGTTCATGCTAGAGTTTTGTTTCCTCCACAGATAT
			Kaspar A2	GAAGGTGGAGTCAACGGATTGAGTTTTGTTTCCTCCACAGATAC
			Kaspar C	ATGCCTTGAATGTGCACCATTCTATGTA
	Sanger seq	Std	Seq F	CTCACGCAGATCACATGACC
			Seq R	GTGGGCGTCTCACATTTTG
rs2811710	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTTATAGCTACTTCAGAAGGCTCAGG
			Kaspar A2	GAAGGTGGAGTCAACGGATTTATAGCTACTTCAGAAGGCTCAGA
			Kaspar C	CTTACCTTGAGAGGACTCTGTGCTA
	Sanger seq	Std	Seq F	AGTCTTGATTTCTGAAAAGGCTA
			Seq R	TTGTTACTTCTACCTTGGTGCA
rs2790457	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTGTGGTGGTACCTGCCTTAGTT
			Kaspar A2	GAAGGTGGAGTCAACGGATTGTGGTGGTACCTGCCTTAGTC
			Kaspar C	CCTTGACCTCCTCAGCTTAAGTGAT
	Sanger seq	Std	Seq F	TGCATTTCAGTGTGGGCTTT
			Seq R	AGTCATAACTCAACAAACCGTC
rs7193541	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTGGACTGGCCAATGGTTCAAG
			Kaspar A2	GAAGGTGGAGTCAACGGATTGCTGGACTGGCCAATGGTTCAA
			Kaspar C	GACTGCTCGTGTTCGCACGTCAT
rs6066835	Kaspar	Std42plus5	Kaspar A1	GAAGGTGACCAAGTTCATGCTGTGCCAGTTCAATCACGCCC
			Kaspar A2	GAAGGTGGAGTCAACGGATTGTGCCAGTTCAATCACGCCT
			Kaspar C	TGCAGGGTCTCAGGCCTGAGAT
	Sanger seq	Std	Seq F	CTGGTGAGGAGGATTCTGGG
			Seq R	GGAACACTTGGACTCTGAGC
rs78311596	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTGGAGATGGTCTATGCTTTTAGGAT
			Kaspar A2	GAAGGTGGAGTCAACGGATTGGAGATGGTCTATGCTTTTAGGAA
			Kaspar C	TAAGGTCACATGGTACGGGA
	Sanger seq	Std	Seq F	CGCTGTGCTCCAACATTCAT
			Seq R	CTGCAGCATTCCAGAGGCT
rs72665486	Kaspar	3stepStd42	Kaspar A1	GAAGGTGACCAAGTTCATGCTTTCCAGTTGAGCTTCGGATAAGAC
			Kaspar A2	GAAGGTGGAGTCAACGGATTTTCCAGTTGAGCTTCGGATAAGAA
			Kaspar C	CACAAGACTGAGATCAAGGTAATCTTCAA
	Sanger seq	Std	Seq F	CACAGTTTGTGGGTTAGGA
			Seq R	TTGAAGACCTAACGGCCCTA
rs1034447	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTAGGAAAACCTCCATGATGCAGTG
			Kaspar A2	GAAGGTGGAGTCAACGGATTACAGGAAAACCTCCATGATGCAGTA
			Kaspar C	GTCTGCACACATCTCCATGTGTATG
	Sanger seq	Std	Seq F	AGTGCCTAAATACATACCACCTT
			Seq R	TCAAGGACAAATGTTCCAGGT
rs17507636	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTGTATAGGGTAGAATCTAATGCTCTGG
			Kaspar A2	GAAGGTGGAGTCAACGGATTGTATAGGGTAGAATCTAATGCTCTGA
			Kaspar C	AGAAGTCTTTCACTGTAGCCATCTGTAT
rs28571765	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTCCTGTAATCCAGCACTTTGAGG
			Kaspar A2	GAAGGTGGAGTCAACGGATTACCTGTAATCCAGCACTTTGAGA

			Kaspar C	TCTCAAACCTCTGGGCTCAAGCAAT
	Sanger seq	Std	Seq F	TCTTACTTCCATCACCCATGT
			Seq R	AGCTATACTATGAATCAAAGCA
rs76601148	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTACAAAATGGGATATCCAATGATTACTACTA
			Kaspar A2	GAAGGTCTGGAGTCAACGGATTCAAATGGGATATCCAATGATTACTACTG
			Kaspar C	CCTGGATAAGATAATCACCGTGTTTCATT
	Sanger seq	Std	Seq F	TGTAGCGACTGGAAAATGGC
			Seq R	CCTGGATAAGATAATCACCGTGT
rs8058578	Kaspar	Std42	Kaspar A1	GAAGGTGACCAAGTTCATGCTTCAACTTCATATTGTTGGTGTTTTCTC
			Kaspar A2	GAAGGTCTGGAGTCAACGGATTCTTCAACTTCATATTGTTGGTGTTTTCTT
			Kaspar C	GGGCGAAAAAGGAAGAGCCAAAGA
rs6071887	Kaspar	Std42plus5	Kaspar A1	GAAGGTGACCAAGTTCATGCTGAACAGAAGAAATAATAAATAGCTGATTGAGT
			Kaspar A2	GAAGGTCTGGAGTCAACGGATTACAGAAGAAATAATAAATAGCTGATTGAGC
			Kaspar C	CGAGATTCTTGAAGAAGGTTGATTACAT
	Sanger seq	Std	Seq F	CCAACCAGCAGAGTTTAGACG
			Seq R	ATGGAGTTCAGCTTGTGGGT

KASPAR conditions

Std42

- Hot Start: 94°C for 15 minutes
- Stage 1: 20 cycles
 - o 94°C for 10 seconds
 - o 57°C for 5 seconds
 - o 72°C for 10 seconds
- Stage 2: 22 cycles
 - o 94°C for 10 seconds
 - o 57°C for 20 seconds
 - o 72°C for 40 seconds

Std42plus5

- Hot Start: 94°C for 15 minutes
- Stage 1: 20 cycles
 - o 94°C for 10 seconds
 - o 57°C for 5 seconds
 - o 72°C for 10 seconds
- Stage 2: 22 cycles
 - o 94°C for 10 seconds
 - o 57°C for 20 seconds
 - o 72°C for 40 seconds
- Stage 3: 5 cycles
 - o 94°C for 10 seconds
 - o 57°C for 1 minute

3StepStd42

- Hot Start: 94°C for 15 minutes
- Stage 1: 42 cycles
 - o 94°C for 10 seconds
 - o 57°C for 20 seconds
 - o 72°C for 40 seconds

Sequencing conditions

Std

- 95°C for 5 minutes
- 25 cycles
 - o 96°C for 30 seconds
 - o 50°C for 15 seconds
 - o 60°C for 1 minute

Supplementary Table 18: Primers and probes and reaction conditions used for this study.

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