

	Percent misclassified cases									
Power	0%	10%	20%							
0.2	1.14	1.15	1.17							
0.5	1.16	1.18	1.21							
0.8	1.19	1.21	1.24							

**Supplementary Figure 1**. Power to detect a variant with misclassifications rates between 0 and 20%. The top graph depicts power as a function of odds ratio for detecting a variant with minor allele frequency 0.1, with misclassification rates of 0%, 10%, and 20% (where the specified fraction of study cases are misclassified controls). The table (bottom) shows odds ratios required to achieve the specified power for a variant with minor allele frequency 0.1, based on the GWAS sample size, for misclassification rates of 0%, 10%, and 20%. To account for misclassification, expected genotype frequencies in study cases were replaced with a mixture of genotype frequencies in true cases and in true controls.

Stage 1, genotyped SNPs only



**Supplementary Figure 2**. Manhattan plot of stage 1 showing associations between genotyped SNPs and BCC. Total stage 1 GWAS analysis included 12,945 cases and 274,252 controls. Loci with smallest  $P < 10^{-6}$  (generated via logistic regression) are labeled with the name of the nearest gene; for clarity, only the 25 most significant loci are labeled. Positions with  $P < 5 \times 10^{-8}$  (genome-wide significance) are shown in red.

Stage 1, genotyped and imputed SNPs



**Supplementary Figure 3**. QQ plot of observed versus expected quantiles for the BCC GWAS stage 1 *P* values (generated via logistic regression), plotted on a log scale. The null hypothesis states that the expected distribution of *P* values is uniform. Here, the observed *P* values follow the null distribution for large *P* values (*P*>0.01) but then diverge for small *P* values. The solid red line has a slope 1 and the dashed red lines represent a 95% confidence interval, assuming the test results are independent. The test statistics in the plot have been adjusted for genomic control.

Stage 1, genotyped SNPs only



**Supplementary Figure 4**. QQ plot of observed versus expected quantiles for stage 1 *P* values, plotted on a log scale, when only genotyped SNPs are included. The dashed line has a slope 1. (Genomic inflation factor,  $\lambda$ = 1.091).



**Supplementary Figure 5**. QQ plot of observed versus expected quantiles for stage 1 *P* values, plotted on a log scale, excluding 28 genome-wide significant loci from stage 1 and their surrounding 500kb interval. The dashed line has a slope 1.



**Supplementary Figure 6**. QQ plot of observed versus expected quantiles for stage 1 *P* values, plotted on a log scale, excluding 37 significant loci (P<10<sup>-6</sup>) from stage 1 and their surrounding 500kb interval. The dashed line has a slope 1.



Stage 2, imputed and genotyped SNPs

**Supplementary Figure 7**. QQ plot of observed versus expected quantiles for stage 2 *P* values, plotted on a log scale. The dashed line has a slope 1. (Genomic inflation factor,  $\lambda$ =1.023).



**Supplementary Figure 8**. Effect sizes estimated in men versus women, for the 28 genome-wide significant loci from stage 1.



**Supplementary Figure 9**. Effect sizes estimated in individuals with melanoma versus individuals without melanoma, for the 28 genome-wide significant loci from stage 1.





**Supplementary Figure 10** 



**Supplementary Figure 10.** Forest plots for each of the 14 novel SNPs reaching genomewide significance for association with BCC (via logistic regression). For all plots, x-axis displays odds ratio (OR) values and gray vertical lines represent an odds ratio (OR) of 1. Each black square indicates OR from the corresponding stage and horizontal gray lines represent 95% CI. The center of each diamond represents the OR for the overall metaanalysis, and the width of each diamond represents the 95% CI.



**Supplementary Figure 11.** Regional association plots for the first 8 of 14 novel, genome-wide significant BCC susceptibility loci are shown. Left to right, beginning from top left: 3p13 (rs2116709, *FOXP1*), 9p22.2 (rs10810657, near *BNC2*), 3q28 (rs191177147, *LPP*), 6p21.32 (rs9275642, *HLA-DQA2*), 7q22.1 (rs73183643, *CUX1*), 8q21.13 (rs11993814, *ZBTB10*), 21q22.3 (rs2776353, *LINCO0111*), 19p13.3.2 (rs10425559, *PLIN3*). Each plot displays -log<sub>10</sub>(*P* value) versus genomic position based on stage 1 logistic regression association testing. The color scale indicates strength of linkage disequilibrium (r<sup>2</sup>) for nearby SNPs, with respect to the index SNP. To preserve detail, results with *P*<10<sup>-100</sup> are set to 10<sup>-100</sup>. The "o" and "+" symbols represent genotyped and imputed SNPs, respectively. Recombination rates, in cM/Mb, are also plotted (navy blue lines). These plots were generated via LocusZoom, using LD data from the March 2012 release of 1000 Genomes data.



**Supplementary Figure 12.** Regional association plots for 6 of 14 novel, genome-wide significant BCC susceptibility loci are shown. Left to right, beginning from top left: 6p21.3 (rs1050529, *HLA-B*), 7p12.3 (rs7776701, *TNS3*), 6p21.3 (rs9267650, *NEU1*), 10q24.3 (rs7907606, *OBFC1*), 6q27 (rs4710154, *MIR3939*), 6p22.3 (rs2294214, *CASC15*). Each plot displays -log<sub>10</sub>(*P* value) versus genomic position based on stage 1 logistic regression association testing.

	Status	n (%)	Male (%)	Age < 31 yr	Age 31-45	Age 46-60	Age > 60
23andMe (Stage 1, n = 287197)	Cases	12945 (4.5)	6700 (52)	42 (0.3)	650 (5)	3194 (25)	9059 (70)
	Controls	274252 (95.5)	148415 (54)	39673 (14)	83162 (30)	74977 (27)	76440 (28)
Harvard (Stage 2, n = 17044)							
Affy	Cases	1777 (24.7)	834 (46.9)	5 (0.3)	500 (28.1)	978 (55.0)	294 (16.5)
	Controls	5411 (75.3)	2385 (44.1)	43 (0.8)	2068 (38.2)	2533 (46.8)	767 (14.2)
Illumina	Cases	1268 (25.6)	368 (29.0)	6 (0.5)	472 (37.2)	678 (53.5)	112 (8.8)
	Controls	3685 (74.4)	1029 (27.9)	33 (0.9)	1681 (45.6)	1692 (45.9)	279 (7.6)
Omni	Cases	1197 (24.4)	447 (37.3)	4 (0.3)	403 (33.7)	645 (53.9)	145 (12.1)
	Controls	3706 (75.6)	1238 (33.4)	34 (0.9)	1671 (45.1)	1651 (44.5)	350 (9.4)
All, Stage 2	Cases	4242 (24.9)	1649 (38.9)	15 (0.4)	1375 (32.4)	2301 (54.2)	551 (13.0)
	Controls	12802 (75.1)	4652 (36.3)	110 (0.9)	5420 (42.3)	5876 (45.9)	1396 (10.9)
Combined meta- analysis (n = 304241)	Cases	17187 (5.6)	8349 (49)	57 (0.3)	2025 (12)	5495 (32)	9610 (56)
, , , ,	Controls	287054 (94.4)	153067 (53)	39783 (14)	88582 (31)	80853 (28)	77836 (27)

### Supplementary Table 1. Gender and age of BCC cases and controls from two-stage GWAS

Counts and percentages for cases and controls (n (%)) are listed above, stratified by stage of GWAS. We also report number and percentage of male subjects, subjects with age < 31 years, subjects with age 30-45 years, subjects with age 45-60 years, and subjects with age > 60 years. Stage 2 cases and controls are further subdivided based on platform used for genotyping.

n = 190	Disease (+)	Disease (-)
Self-report (+) Self-report (-)	42 3	1 144
	Sensitivity = 9 Specificity = 9	93% 99%

# Supplementary Table 2. Sensitivity and specificity of self-report data with respect to BCC diagnosis

Within table, from left to right, are counts for true positives, false positives, false negatives, and true negatives. Data from 190 randomly selected patients at Stanford outpatient clinics.

Locus	Nearby genes	Imputed SNP	Imputed position	Imputation <i>P</i> -value	Genotyped SNP	Genotyped position	Genotyped <i>P</i> -value
6p25.3	IRF4	rs12203592	396321	7.9E-128	rs12210050	475489	1.6E-40
16q24.3	MC1R	rs1805007	89986117	3.5E-48	rs1805007	89986117	1.4E-45
20p13	TGM3	rs214785	2283457	4.4E-26	rs214796	2286343	3.8E-26
5p13.2	SLC45A2	rs35407	33946571	5.8E-26	rs16891982	33951693	3.2E-36
20q11.22	RALY	rs6059655	32665748	2.5E-22	rs910873	33171772	2.4E-13
1p36.13	RCC2	rs57142672	17744536	9.3E-21	rs6688886	17753639	1.9E-20
10p14	GATA3	rs73635312	8935998	8.5E-20	rs290903	8936140	8.5E-16
11q14.3	TYR	rs1126809	89017961	7.7E-19	rs1393350	89011046	1.1E-16
5p15.33	CLPTM1L	rs421284	1325590	2.0E-17	rs401681	1322087	3.4E-16
2q33.1	ALS2CR12/CASP8	rs2080303	202165206	1.4E-14	rs10931936	202143928	1.2E-13
1q42.13	RHOU	rs61824911	228979737	4.4E-14	rs71074	229003739	4.5E-13
3p13	FOXP1/EIF4E3	rs2116709	71621669	8.7E-14	rs9828619	71626123	1.4E-13
15q13.1	OCA2/HERC2	rs12916300	28410491	1.5E-13	rs16950987	28526228	6.0E-07
8q22.2	RGS22	rs141115006	101005020	2.4E-13	rs7006527	101024505	1.4E-11
9p22.2	BNC2	rs10810657	16884586	4.9E-13	rs2153271	16864521	5.3E-11
8q21.11	ZFHX4	rs10093547	77474545	7.4E-12	rs17431544	77467488	1.4E-11
3q28	LPP	rs191177147	188092546	2.2E-11	rs9851967	188087628	3.5E-10
6p21.32	HLA-DQB1/DQA2	rs9275642	32684935	7.7E-11	rs4947342	32653070	3.3E-09
9p21.3	CDKN2B	rs7874604	22054690	1.8E-10	rs7044859	22018781	3.1E-09
12q13.13	KRT5	rs11170164	52913668	2.7E-10	rs11170164	52913668	1.9E-10
7q22.1	CUX1	rs73183643	101418653	4.6E-10	rs11977362	101381523	3.3E-06
8q21.13	ZBTB10	rs11993814	81389791	1.4E-09	rs7830160	81381871	1.9E-09
21q22.3	LINC00111	rs2776353	43089017	2.3E-09	rs2776348	43086944	3.6E-08
13q34	ATP11A	rs1765871	113533651	4.9E-09	rs1046793	113539894	1.8E-08
19p13.3	TICAM1/PLIN3	rs10425559	4837487	1.5E-08	rs6510827	4830628	7.5E-08
6p21.33	HLA-B	rs1050529	31324615	1.8E-08	rs2524043	31257012	4.5E-05
7p12.3	TNS3	rs7776701	47442633	2.1E-08	rs834602	47446215	1.9E-06
22q12.1	TTC28	rs78097823	28648766	4.7E-08	rs9625433	28620996	4.5E-05

Supplementary Table 3. Association of 28 imputed and genotyped index SNPs from stage 1 with BCC

For each index SNP from stage 1, we show the best association test result for a nearby SNP that was genotyped on our V3 and/or V4 arrays. "Imputed *P*-value" refers to the association (via logistic regression) between the stage 1 imputed index SNPs and BCC. "Genotyped *P*-value" refers to the association (via logistic regression) between stage 1 genotyped SNPs and BCC.

					Stage 2						Meta-analysis					
SNP	Region	Gene	Maj/ min	Ρ	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	P <sub>het</sub>	ľ	
rs12203592	6p25.3	IRF4	C/T	3.1×10 <sup>-10</sup>	1.31	(1.21, 1.43)	-	1.00	1.00	1.00	1.5×10 <sup>-151</sup>	1.48	(1.44, 1.53)	0.00	88%	
rs11170164	12q13.13	KRT5	C/T	3.4×10 <sup>-5</sup>	1.30	(1.15, 1.47)	-	0.85	0.99	0.92	1.2×10 <sup>-14</sup>	1.19	(1.14, 1.24)	0.13	56%	
rs10425559	19p13.3	PLIN3	G/A	8.4×10 <sup>-1</sup>	0.99	(0.93, 1.06)	-	0.88	1.00	0.94	2.8×10 <sup>-8</sup>	0.93	(0.91, 0.96)	0.05	74%	
rs9267650	6p21.33	NEU1	A/T	3.6×10 <sup>-1</sup>	1.08	(0.92, 1.27)	-	0.86	0.98	0.92	1.5×10 <sup>-8</sup>	1.17	(1.11, 1.24)	0.29	9%	
rs12210050	6p25.3	EXOC2	C/T	3.3×10 <sup>-5</sup>	1.17	(1.09, 1.27)	1.00	1.00	-	1.00	1.0×10 <sup>-49</sup>	1.25	(1.21, 1.29)	0.07	69%	

Supplementary Table 4. Meta-analysis for 5 SNPs using subsets with high imputation quality in stage 2

For each SNP, we report rsID, genetic locus, gene, major and minor alleles, *P* value for stage 2, odds ratio (for stage 2 overall, along with 95% CI), average imputation  $r^2$  (Affy Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and *P*<sub>het</sub> and *I*<sup>2</sup> for meta-analysis. *P* values generated via logistic regression.

Supplementary Table 5. Heterogeneity analysis results between stage 2 genotyping and imputation data	

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					Imputation						Genotyping						
SNP	Region	Gene	Maj/ Min	MAF	Р	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	MAF	Р	OR	95% CI	P <sub>het</sub>	/2
rs12916300	15q13.1	HERC2	T/C	0.35	5.0×10 <sup>-4</sup>	0.84	(0.76, 0.93)	0.47	0.31	0.11	0.30	0.21	3.0×10 <sup>-1</sup>	0.91	(0.75, 1.09)	0.47	0%
rs35407	5p13.2	SLC45A2	G/A	0.05	7.4×10 <sup>-5</sup>	0.71	(0.61, 0.84)	0.60	0.58	0.33	0.50	0.02	8.2×10 <sup>-4</sup>	0.43	(0.26, 0.71)	0.06	72%

For each SNP, we report rsID, genetic locus, gene, major and minor alleles, minor allele frequency (MAF), *P* value and odds ratio (for stage 2 imputation, along with 95% CI), average imputation  $r^2$  (Affy, Illumina, Omni, stage 2 overall), MAF for genotyping, *P* value and odds ratio (for stage 2 genotyping, along with 95% CI), and *P*<sub>het</sub> and *I*<sup>2</sup>. *P* values generated via logistic regression.

		Age interval								
SNP	Gene	(0,57]	(57,65]	(65,72]	(72,72+]	P-value				
rs12203592	IRF4	0.509	0.395	0.434	0.275	1.30E-05				
rs1805007	MC1R	0.406	0.345	0.392	0.200	0.0063				
rs214785	TGM3	-0.246	-0.207	-0.118	-0.154	0.044				
rs35407	SLC45A2	0.725	0.478	0.650	0.255	0.0065				
rs6059655	RALY	-0.264	-0.284	-0.197	-0.187	0.061				
rs57142672	RCC2	0.140	0.140	0.134	0.114	0.8				
rs73635312	GATA3	0.257	0.223	0.111	0.145	0.038				
rs1126809	TYR	-0.133	-0.175	-0.102	-0.114	0.18				
rs421284	CLPTM1L	0.119	0.118	0.131	0.091	0.88				
rs2080303	ALS2CR12/CASP8	0.130	0.129	0.118	0.098	0.99				
rs61824911	RHOU	0.189	0.092	0.072	0.126	0.067				
rs2116709	FOXP1/EIF4E3	-0.115	-0.105	-0.110	-0.114	0.99				
rs12916300	OCA2/HERC2	0.185	0.127	0.114	0.106	0.25				
rs141115006	RGS22	-0.127	-0.123	-0.132	-0.182	0.66				
rs10810657	BNC2	-0.146	-0.078	-0.105	-0.082	0.24				
rs10093547	ZFHX4	0.171	0.159	0.269	0.198	0.5				
rs191177147	LPP	0.123	0.057	0.122	0.106	0.3				
rs9275642	HLA-DQB1/DQA2	-0.125	-0.116	-0.097	-0.123	0.9				
rs7874604	CDKN2B	0.078	0.076	0.111	0.115	0.78				
rs11170164	KRT5	0.225	0.143	0.198	0.049	0.079				
rs73183643	CUX1	0.141	0.128	0.061	0.084	0.42				
rs11993814	ZBTB10	-0.060	-0.096	-0.104	-0.115	0.54				
rs2776353	LINC00111	-0.103	-0.100	-0.097	-0.052	0.6				
rs1765871	ATP11A	-0.074	-0.049	-0.123	-0.072	0.32				
rs10425559	TICAM1/PLIN3	0.094	0.086	0.042	0.091	0.36				
rs1050529	HLA-B	-0.124	-0.084	-0.068	-0.131	0.35				
rs7776701	TNS3	-0.132	-0.086	-0.057	-0.033	0.048				
rs78097823	TTC28	0.078	0.289	0.180	0.185	0.11				

## Supplementary Table 6. Effect sizes of 28 index SNPs in stage 1 stratified by age

The table shows effect sizes (for each SNP reaching genome-wide significance in stage 1) computed in each age interval, as well as the *P* value for the test of genotype interaction with age interval. Stage 1 cohort was divided into four age intervals with similar effective sample sizes based on case and control sample counts. For all these association tests, the same covariates were used as in stage 1: age, sex, and five principal components. Thus, association tests within a specific age interval were still adjusted for age as a continuous covariate. Mean age for the 274,252 controls was 48.9 (SD 16.2); for the 12,945 cases, mean age was 65.5 (SD 11.5).

SNP	Gene	M <sub>effect</sub>	M <sub>SE</sub>	M <sub>P</sub>	F <sub>effect</sub>	F <sub>SE</sub>	F <sub>P</sub>	Р
rs12203592	IRF4	0.455	0.022	7.0E-88	0.366	0.023	6.0E-55	0.040
rs1805007	MC1R	0.374	0.031	4.1E-32	0.316	0.031	1.8E-22	0.286
rs214785	TGM3	-0.167	0.023	1.2E-12	-0.199	0.023	4.3E-17	0.213
rs35407	SLC45A2	0.505	0.071	1.9E-14	0.541	0.074	1.9E-15	0.751
rs6059655	RALY	-0.279	0.032	1.3E-17	-0.194	0.032	4.4E-09	0.088
rs57142672	RCC2	0.143	0.019	1.6E-13	0.125	0.020	2.1E-10	0.612
rs73635312	GATA3	0.227	0.028	2.2E-16	0.141	0.028	3.3E-07	0.047
rs1126809	TYR	-0.128	0.020	2.8E-10	-0.140	0.021	1.5E-11	0.521
rs421284	CLPTM1L	0.116	0.019	4.7E-10	0.120	0.019	3.0E-10	0.783
rs2080303	ALS2CR12/CASP8	0.135	0.021	1.1E-10	0.105	0.021	8.2E-07	0.425
rs61824911	RHOU	0.139	0.021	9.1E-11	0.102	0.022	3.9E-06	0.300
rs2116709	FOXP1/EIF4E3	-0.133	0.020	2.6E-11	-0.087	0.020	1.7E-05	0.136
rs12916300	OCA2/HERC2	0.136	0.025	2.2E-08	0.133	0.025	9.3E-08	0.761
rs141115006	RGS22	-0.140	0.026	7.7E-08	-0.144	0.027	6.9E-08	0.819
rs10810657	BNC2	-0.104	0.019	4.8E-08	-0.102	0.020	2.0E-07	0.986
rs10093547	ZFHX4	0.217	0.041	7.8E-08	0.191	0.042	2.8E-06	0.755
rs191177147	LPP	0.098	0.021	3.4E-06	0.111	0.021	2.2E-07	0.517
rs9275642	HLA-DQB1/DQA2	-0.137	0.025	2.7E-08	-0.100	0.025	5.7E-05	0.346
rs7874604	CDKN2B	0.088	0.020	1.5E-05	0.106	0.021	3.7E-07	0.591
rs11170164	KRT5	0.186	0.033	3.4E-08	0.133	0.034	1.4E-04	0.303
rs73183643	CUX1	0.094	0.023	3.2E-05	0.115	0.023	5.1E-07	0.391
rs11993814	ZBTB10	-0.110	0.022	3.5E-07	-0.083	0.022	1.6E-04	0.413
rs2776353	LINC00111	-0.084	0.020	3.8E-05	-0.097	0.021	3.1E-06	0.623
rs1765871	ATP11A	-0.082	0.019	1.0E-05	-0.080	0.019	2.6E-05	0.981
rs10425559	TICAM1/PLIN3	0.093	0.019	1.1E-06	0.067	0.020	6.3E-04	0.458
rs1050529	HLA-B	-0.095	0.025	1.6E-04	-0.116	0.026	6.4E-06	0.486
rs7776701	TNS3	-0.074	0.019	7.2E-05	-0.081	0.019	2.2E-05	0.711
rs78097823	TTC28	0.212	0.045	4.9E-06	0.166	0.046	4.0E-04	0.466

Supplementary Table 7. Gender-based logistic regression results for 28 index SNPs in stage 1

The table shows results from logistic regression models (for each SNP reaching genome-wide significance in stage 1) fit separately in male and female subsets of the stage 1 cohort, and a *P*-value from a likelihood ratio test for adding a gender by genotype interaction to the full model. "M" stands for male and "F" for female. The subscripts "SE" and "P" stand for "standard error" and "P-value", respectively. Of the 274,252 controls, 54% were male; of the 12,945 cases, 52% were male.

			Mai/	MAF (avg	Stage	1	Stage 2		Meta-ana	lysis	Prior Studies		
SNP	Region	Gene	min	imputation $r^2$ )	P	OR	Р	OR	Р	OR	Р	OR	Ref
rs12203592	6p25.3	IRF4*	C/T	0.17 (0.99)	1.5×10 <sup>-138</sup>	1.51	1.5×10 <sup>-11</sup>	1.31	2.4×10 <sup>-152</sup>	1.48	1.6×10 <sup>-3</sup>	1.32	а
rs1805007	16q24.3	MC1R	C/T	0.07 (1.0)	4.0×10 <sup>-52</sup>	1.41	1.9×10 <sup>-9</sup>	1.34	2.5×10 <sup>-63</sup>	1.40	4.3×10 <sup>-17</sup>	1.55	4
rs12210050	6p25.3	EXOC2	C/T	0.17 (0.99)	3.3×10 <sup>-45</sup>	1.27	4.5×10 <sup>-7</sup>	1.19	1.0×10 <sup>-51</sup>	1.25	9.9×10 <sup>-10</sup>	1.24	4
rs214785	20p13	TGM3	T/C	0.18 (0.99)	3.7×10 <sup>-28</sup>	1.2	1.2×10 <sup>-5</sup>	1.15	7.9×10 <sup>-33</sup>	1.19	5.5×10 <sup>-17</sup>	1.29	6
rs35407	5p13.2	SLC45A2*	G/A	0.04 (0.98)	5.0×10 <sup>-28</sup>	0.6	7.4×10 <sup>-5</sup>	0.71	5.2×10 <sup>-27</sup>	0.63	1.6×10 <sup>-12</sup>	0.51	b
rs6059655	20q11.22	RALY*	G/A	0.07 (0.99)	4.2×10 <sup>-24</sup>	1.26	2.8×10 <sup>-3</sup>	1.15	2.5×10 <sup>-26</sup>	1.24	1.2×10 <sup>-6</sup>	1.35	C**
rs57142672	1p36.13	RCC2	A/G	0.34 (0.99)	2.2×10 <sup>-22</sup>	1.14	3.4×10 <sup>-3</sup>	1.08	1.0×10 <sup>-23</sup>	1.13	4.4×10 <sup>-12</sup>	1.28	2
rs73635312	10p14	GATA3	G/A	0.14 (1.0)	2.4×10 <sup>-21</sup>	0.83	2.3×10 <sup>-4</sup>	0.87	2.8×10 <sup>-23</sup>	0.84	2.4×10 <sup>-16</sup>	0.74	7
rs1126809	11q14.3	TYR*	G/A	0.28 (0.99)	2.6×10 <sup>-20</sup>	1.14	1.3×10 <sup>-1</sup>	1.04	2.5×10 <sup>-19</sup>	1.12	6.1×10 <sup>-4</sup>	1.14	С
rs421284	5p15.33	CLPTM1L	T/C	0.44 (0.99)	9.1×10 <sup>-19</sup>	0.89	2.8×10 <sup>-2</sup>	0.94	1.1×10 <sup>-18</sup>	0.90	3.7×10 <sup>-12</sup>	0.80	3
rs2080303	2q33.1	ALS2CR12	C/T	0.32 (0.89)	1.1×10 <sup>-15</sup>	1.13	1.9×10 <sup>-4</sup>	1.11	7.4×10 <sup>-19</sup>	1.13	1.5×10 <sup>-9</sup>	1.15	7
rs61824911	1q42.13	RHOU	A/G	0.28 (0.90)	3.7×10 <sup>-15</sup>	1.13	1.4×10 <sup>-1</sup>	1.05	1.1×10 <sup>-14</sup>	1.11	5.9×10 <sup>-12</sup>	1.28	2
rs12916300	15q13.1	OCA2*	T/C	0.29 (0.79)	1.4×10 <sup>-14</sup>	0.87	5.0×10 <sup>-4</sup>	0.84	8.2×10 <sup>-17</sup>	0.87	1×10 <sup>-2</sup>	0.70	а
rs141115006	8q22.2	RGS22	C/T	0.17 (0.95)	2.3×10 <sup>-14</sup>	0.87	5.1×10 <sup>-3</sup>	0.91	2.0×10 <sup>-15</sup>	0.88	8.7×10 <sup>-13</sup>	0.77	6
rs10093547	8q21.11	ZFHX4	T/G	0.06 (0.95)	9.6×10 <sup>-13</sup>	0.82	2.7×10 <sup>-4</sup>	0.82	4.6×10 <sup>-15</sup>	0.82	3.5×10 <sup>-12</sup>	0.70	7
rs7874604	9p21.3	CDKN2B	T/C	0.46 (0.84)	3.0×10 <sup>-11</sup>	0.91	3.8×10 <sup>-3</sup>	0.92	4.5×10 <sup>-13</sup>	0.91	3.0×10 <sup>-10</sup>	0.83	6
rs11170164	12q13.13	KRT5	C/T	0.08 (1.0)	4.7×10 <sup>-11</sup>	1.17	3.7×10 <sup>-6</sup>	1.26	1.1×10 <sup>-15</sup>	1.19	9.3×10 <sup>-9</sup>	1.29	7
rs78378222	17p13.1	TP53	T/G	0.01 (0.87)	3.5×10 <sup>-8</sup>	1.39	4.0×10 <sup>-3</sup>	1.54	1.8×10 <sup>-10</sup>	1.41	2.2×10 <sup>-20</sup>	2.16	5
rs157935	7q32.3	KLF14	T/G	0.29 (1.0)	6.5×10 <sup>-5</sup>	0.94	2.3×10 <sup>-2</sup>	0.94	4.5×10 <sup>-6</sup>	0.94	8.5×10 <sup>-11</sup>	0.81	6
rs57244888	2p24.3	MYCN	T/C	0.12 (0.92)	6.6×10 <sup>-5</sup>	0.92	4.4×10 <sup>-1</sup>	0.97	1.2×10 <sup>-4</sup>	0.93	4.7×10 <sup>-12</sup>	0.76	7
rs7335046	13q32.3	UBAC2	C/G	0.12 (0.99)	5.6×10 <sup>-1</sup>	1.01	2.5×10 <sup>-4</sup>	1.15	2.6×10 <sup>-2</sup>	1.04	2.9×10 <sup>-8</sup>	1.26	4

applementally rable of hephtation of zo previously committee and o previously reported bee associated re	Supplementary Table 8.	Replication of 16	previously	y confirmed and 5	previously r	reported BCC-	-associated lo
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16 loci previously confirmed as associated with BCC via prior GWAS ( $P < 5 \times 10^{-8}$ ) are listed, 12 of which independently reached genome-wide significance ( $P < 5 \times 10^{-8}$ ) in this study. 5 additional BCC loci, previously reported via candidate gene studies, are indicated via asterisks; these loci are pigmentation-related, and all 5 reached genome-wide significance in this study. We report the most significant SNP at each locus based on stage 1 data. Additionally, we report major allele, minor allele frequency (MAF) as calculated from stage 1 data, average imputation  $r^2$  (a measure of imputation quality) for stage 1, and odds ratio (OR) with *P* value for each stage, calculated with respect to the minor allele via logistic regression. The right-most 3 columns list *P* value and OR from prior publications for each locus, along with corresponding reference number ("Ref") as listed in main text. Statistics for effect heterogeneity ( $P_{het}$  and  $I^2$ ) are included in Supplementary Table 9.

a. Han, J. et al. A germline variant in the interferon regulatory factor 4 gene as a novel skin cancer risk locus. Cancer Res. 71, 1533 1539 (2011).

b. Stacey, S. N. et al. New common variants affecting susceptibility to basal cell carcinoma. Nat. Genet. 41, 909–914 (2009).

c. Gudbjartsson, D. F. et al. ASIP and TYR pigmentation variants associate with cutaneous melanoma and basal cell carcinoma. Nat. Genet. 40, 886–891 (2008).

\*\*This publication reports P value and OR with respect to a two-SNP haplotype, rs1015362[G] and rs4911414[T], at the RALY-ASIP locus

			Stage 1			Stage 2					Meta-analysis	
SNP	Gene	Maj/ min	MAF	avg r <sup>2</sup>	min r <sup>2</sup>	MAF	Affy r <sup>2</sup>	lllum r <sup>2</sup>	Omni r <sup>2</sup>	Overall avg r <sup>2</sup>	P <sub>het</sub>	
10000500		o /=	0.47			0.47			4.00	0.77		
rs12203592	IRF4*	C/1	0.17	0.99	0.98	0.17	0.32	1.00	1.00	0.77	0.00	91%
rs1805007	MC1R	C/T	0.07	1.00	1.00	0.07	1.00	0.82	0.88	0.90	0.32	0%
rs12210050	EXOC2	C/T	0.17	0.99	0.98	0.17	1.00	1.00	0.54	0.84	0.13	56%
rs214785	TGM3	T/C	0.18	0.99	0.98	0.18	1.00	0.99	0.99	0.99	0.27	17%
rs35407	SLC45A2*	G/A	0.04	0.98	0.88	0.05	0.60	0.58	0.33	0.50	0.06	71%
rs6059655	RALY*	G/A	0.07	0.99	0.98	0.08	0.97	0.98	0.99	0.98	0.07	70%
rs57142672	RCC2	A/G	0.34	0.99	0.98	0.35	0.98	0.98	0.98	0.98	0.06	71%
rs73635312	GATA3	G/A	0.14	1.00	1.00	0.14	0.97	1.00	0.99	0.98	0.27	16%
rs1126809	TYR*	G/A	0.28	0.99	0.98	0.27	0.98	0.97	0.96	0.97	0.00	88%
rs421284	CLPTM1L	T/C	0.44	0.99	0.99	0.44	0.99	0.98	0.99	0.99	0.03	78%
rs2080303	ALS2CR12	C/T	0.32	0.89	0.88	0.32	0.88	0.88	0.88	0.88	0.71	0%
rs61824911	RHOU	A/G	0.28	0.90	0.89	0.27	0.89	0.88	0.88	0.88	0.03	80%
rs12916300	OCA2*	T/C	0.29	0.79	0.75	0.35	0.47	0.31	0.11	0.30	0.40	0%
rs141115006	RGS22	C/T	0.17	0.95	0.92	0.17	0.98	0.92	0.96	0.96	0.27	17%
rs10093547	ZFHX4	T/G	0.06	0.95	0.93	0.07	0.93	0.93	0.94	0.93	0.95	0%
rs7874604	CDKN2B	T/C	0.46	0.84	0.82	0.45	0.84	0.81	0.82	0.82	0.63	0%
rs11170164	KRT5	C/T	0.08	1.00	0.99	0.08	0.73	0.85	0.99	0.86	0.19	42%
rs78378222	TP53	T/G	0.01	0.87	0.82	0.01	0.79	0.79	0.79	0.79	0.53	0%
rs157935	KLF14	T/G	0.29	1.00	1.00	0.30	0.97	1.00	1.00	0.99	0.85	0%
rs57244888	MYCN	T/C	0.12	0.92	0.90	0.12	0.99	0.88	0.91	0.93	0.23	30%
rs7335046	UBAC2	C/G	0.12	0.99	0.97	0.12	0.98	0.98	0.99	0.98	0.00	89%

Supplementary Table 9. Imputation and effect heterogeneity statistics for SNPs at previously reported loci

These loci are all previously reported in prior BCC GWAS publications (16 loci) or candidate gene studies (5 loci, asterisks). We report genetic context, major and minor alleles, stage 1 minor allele frequency (MAF), stage 1 average imputation  $r^2$  (avg  $r^2$ ), stage 1 minimum imputation  $r^2$ , stage 2 MAF, stage 2 average imputation  $r^2$  for each genotyping platform (Affy, Illumina, Omni) and overall, and *P* value (*P*<sub>het</sub>) and *I*<sup>2</sup> for effect heterogeneity pertaining to meta-analysis of combined stage 1-stage 2 data.

Supplementary Table 10. Meta-analysis for rs12913832 and rs1800407

					St	tage 1					Sta	age 2						Meta-an	nalysis		
SNP	Region	Gene	Maj/ min	MAF	Р	OR	95% CI	Avg r <sup>2</sup>	MAF	Р	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	<b>P</b> <sub>het</sub>	I <sup>2</sup>
rs12913832	15q13.1	OCA2/ HERC2	G/A	0.28	1.0×10 <sup>-11</sup>	0.87	(0.84, 0.91)	1.00	0.26	6.2×10 <sup>-4</sup>	0.89	(0.83 <i>,</i> 0.95)	0.99	1.00	NA	1.00	4.4×10 <sup>-14</sup>	0.88	(0.85 <i>,</i> 0.91)	0.73	0%
rs1800407	15q13.1	OCA2	C/T	0.07	4.1×10 <sup>-8</sup>	1.14	(1.09, 1.20)	1.00	0.08	2.0×10 <sup>-1</sup>	1.12	(0.94 <i>,</i> 1.34)	NA	NA	1.00	1.00	3.2×10 <sup>-8</sup>	1.14	(1.09 <i>,</i> 1.19)	0.46	0%

*OCA2/HERC2* rs12913832 is in modest linkage disequilibrium with rs12916300 ( $r^2$ =0.66, D'=0.93). 15q13.1 (rs1800407; *OCA2* R419Q) is a previously confirmed pigmentation-related locus. For each SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 1 minor allele frequency (MAF), P value for stage 1, odds ratio (for stage 1 overall, plus 95% CI), average imputation  $r^2$  (for stage 2, P value for stage 2, odds ratio (for stage 2 overall, plus 95% CI), average imputation  $r^2$  (for Affy, Illumina, Omni, stage 2 overall), P value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and  $P_{het}$  and  $I^2$  for meta-analysis. P values generated via logistic regression.

Supplementar	y Table 11.	. Meta-anal	ysis for	rs16891982
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					S	Stage 1					S	tage 2						Meta-a	nalysis		
SNP	Region	Gene	Maj/ min	MAF	Р	OR	95% CI	Avg r <sup>2</sup>	MAF	Р	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	P <sub>het</sub>	ľ
rs16891982	5p13.2	SLC4 5A2	G/C	0.05	1.7×10 <sup>-25</sup>	0.59	(0.53 <i>,</i> 0.65)	1.00	0.07	1.2×10 <sup>-4</sup>	0.75	(0.64 <i>,</i> 0.87)	0.56	0.58	0.32	0.49	5.7×10 <sup>-24</sup>	0.64	(0.58 <i>,</i> 0.70)	0.01	84%

rs16891982 (Phe374Leu) in *SLC45A2* is non-synonymous exonic SNP. rs16891982 is in modest LD with rs35407 ( $r^2$ =0.33, D'=1). For each SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 1 minor allele frequency (MAF), *P* value for stage 1, odds ratio (for stage 1 overall, plus 95% CI), average imputation  $r^2$  (for stage 1), MAF for stage 2, *P* value for stage 2, odds ratio (for stage 2 overall, plus 95% CI), average imputation  $r^2$  (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and  $P_{het}$  and  $l^2$  for meta-analysis. *P* values generated via logistic regression.

Supplementary Table 12. Mutually adjusted analysis for rs12203592 and rs12210050 in region 6p25.3 using subsets with high imputation quality in stage 2

					Indepen	ident analysi	s results <sup>c</sup>	Mutually adju	isted ana	lysis results <sup>c</sup>
SNP	Region	Gene	Maj/ Min	r²	P	OR	95% CI	Р	OR	95% CI
rs12203592ª	6p25.3	IRF4	C/T	0.997	4.3×10 <sup>-4</sup>	1.24	(1.10,1.40)	9.9×10 <sup>-3</sup>	1.21	(1.05,1.38)
rs12210050 <sup>b</sup>	6p25.3	EXOC2	C/T	0.996	1.1×10 <sup>-2</sup>	1.17	(1.04,1.32)	3.7×10 <sup>-1</sup>	1.07	(0.93,1.22)

The second signal in the 6p25.3 region peaked at rs12210050, located near *EXOC2*; this SNP was previously reported to be associated with BCC risk and replicated in our study. This SNP is not in strong linkage disequilibrium with the top SNP in 6p25.3, rs12203592 ( $r^2$ =0.26, D'= 0.51). We performed a conditional analysis mutually adjusting for both SNPs in stage 2, using a high-quality Illumina dataset ( $R^2$  > 0.99 for both SNPs). The results demonstrate that rs12203592 remained significant but rs12210050 did not. For each SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 2 average imputation  $r^2$ , and *P* value and OR (along with 95% CI) for independent analysis results and mutually adjusted analysis results, generated via logistic regression.

a. Mutually adjusted analysis adjusted by SNP rs12210050

b. Mutually adjusted analysis adjusted by top SNP rs12203592

c. Analysis was generated using subsets with high imputation quality in stage 2

				Stage 1				Stage 2			Meta-a	nalysis
SNP	Gene	Maj/ min	MAF	avg r <sup>2</sup>	min r <sup>2</sup>	MAF	Affy r <sup>2</sup>	lllum r <sup>2</sup>	Omni r <sup>2</sup>	Overall avg r <sup>2</sup>	P <sub>het</sub>	<i>I</i> <sup>2</sup> (%)
rs2116709	FOXP1	A/T	0.40	0.91	0.90	0.41	0.90	0.89	0.90	0.89	0.58	0%
rs10810657	BNC2	A/T	0.41	0.98	0.97	0.40	0.98	0.97	0.97	0.97	0.89	0%
rs191177147	LPP	G/T	0.39	0.80	0.79	0.39	0.75	0.78	0.78	0.77	0.81	0%
rs9275642	HLA-DQA2	C/T	0.21	0.89	0.81	0.22 <sup>ª</sup>	а	а	а	а	1.00	0%
rs73183643	CUX1	G/A	0.24	0.96	0.93	0.24	0.98	0.94	0.92	0.94	0.76	0%
rs11993814	ZBTB10	C/T	0.26	1.00	0.98	0.25	0.98	0.99	1.00	0.99	0.26	21%
rs2776353	LINC00111	A/T	0.32	0.96	0.93	0.32	0.91	0.91	0.96	0.93	0.85	0%
rs10425559	PLIN3	G/A	0.40	0.97	0.89	0.40	0.28	0.88	1.00	0.72	0.05	74%
rs1050529	HLA-B	C/T	0.25	0.71	0.69	0.19 <sup>b</sup>	b	b	b	b	0.91	0%
rs7776701	TNS3	C/T	0.48	0.98	0.97	0.48	0.98	0.98	0.98	0.98	0.04	77%
rs9267650	NEU1	A/T	0.05	0.98	0.98	0.05	0.66	0.86	0.98	0.83	0.26	23%
rs7907606	OBFC1	T/G	0.17	0.96	0.92	0.17	0.97	0.92	0.97	0.96	0.63	0%
rs4710154	MIR3939	A/T	0.32	0.93	0.93	0.31	0.91	0.92	0.92	0.91	0.53	0%
rs2294214	CASC15	A/C	0.32	0.95	0.93	0.30	0.82	0.95	0.74	0.84	0.07	69%

Supplementary Table 13.	Imputation and effe	ect heterogeneity	statistics for	14 novel	genome-wide
significant SNPs					

SNPs that met genome-wide significance ( $P < 5 \times 10^{-8}$ , via logistic regression) in stage 1 and/or overall meta-analysis are listed. Additionally, we report genetic context, major and minor alleles, stage 1 minor allele frequency (MAF), stage 1 average imputation  $r^2$  (avg  $r^2$ ), stage 1 minimum imputation  $r^2$ , stage 2 MAF, stage 2 average imputation  $r^2$  for each genotyping platform (Affy, Illumina, Omni) and overall, and P value ( $P_{her}$ ) and  $I^2$  for effect heterogeneity pertaining to meta-analysis of combined stage 1-stage 2 data.

a. Due to low imputation quality in stage 2 ( $r^2$  <0.3), genotyping results are utilized for this result.

b. Due to low imputation quality in stage 2 ( $r^2$  <0.3), genotyping results are utilized for this result. rs1050529 is located at HLA region, harboring high genomic diversity. rs9266772 is used as proxy SNP for rs1050529 ( $r^2$ =0.569,D'=0.771).

SNP	Region	Gene	Maj/ min	β	SE	P value	Adj βª	Adj SE <sup>a</sup>	Adj <i>P</i> value <sup>ª</sup>	Note
rs12203592	6p25.3	IRF4	C/T	0.269	0.04	1.5E-11	0.281	0.045	3.1E-10	PC
rs1805007	16q24.3	MC1R	C/T	0.291	0.049	1.9E-09	0.223	0.054	3.0E-05	PC
rs12210050	6p25.3	EXOC2	C/T	0.177	0.035	4.5E-07	0.158	0.038	3.7E-05	PC
rs214785	20p13	TGM3	T/C	-0.143	0.033	1.2E-05	0.131	0.035	2.1E-04	PC
rs35407	5p13.2	SLC45A2	G/A	-0.336	0.085	7.4E-05	-0.189	0.09	3.5E-02	PC
rs6059655	20q11.22	RALY	G/A	0.14	0.047	2.8E-03	0.035	0.051	5.0E-01	PC
rs57142672	1p36.13	RCC2	A/G	-0.078	0.027	3.4E-03	0.1	0.029	4.8E-04	PC
rs73635312	10p14	GATA3	G/A	-0.139	0.038	2.3E-04	-0.158	0.041	1.1E-04	PC
rs1126809	11q14.3	TYR	G/A	0.043	0.029	1.3E-01	0.018	0.031	5.5E-01	PC
rs421284	5p15.33	CLPTM1L	T/C	0.057	0.026	2.8E-02	-0.072	0.028	1.0E-02	PC
rs2080303	2q33.1	ALS2CR12	C/T	0.108	0.029	1.9E-04	0.123	0.031	8.3E-05	PC
rs61824911	1q42.13	RHOU	A/G	-0.045	0.031	1.4E-01	0.056	0.033	9.4E-02	PC
rs12916300	15q13.1	OCA2	T/C	0.18	0.052	5.0E-04	-0.102	0.054	5.6E-02	PC
rs141115006	8q22.2	RGS22	C/T	-0.098	0.035	5.1E-03	-0.079	0.038	3.7E-02	PC
rs10093547	8q21.11	ZFHX4	T/G	0.2	0.055	2.7E-04	-0.254	0.06	2.3E-05	PC
rs7874604	9p21.3	CDKN2B	T/C	0.081	0.028	3.8E-03	-0.053	0.03	8.1E-02	PC
rs11170164	12q13.13	KRT5	C/T	0.232	0.05	3.7E-06	0.224	0.054	3.7E-05	PC
rs78378222	17p13.1	TP53	T/G	-0.433	0.15	4.0E-03	0.36	0.14	1.0E-02	PC
rs157935	7q32.3	KLF14	T/G	0.064	0.028	2.3E-02	-0.066	0.031	3.0E-02	PC
rs57244888	2p24.3	MYCN	T/C	0.031	0.041	4.4E-01	-0.009	0.044	8.4E-01	PC
rs7335046	13q32.3	UBAC2	C/G	-0.139	0.038	2.5E-04	0.134	0.041	9.5E-04	PC
rs2116709	3p13	FOXP1	A/T	0.094	0.027	6.1E-04	-0.094	0.03	1.4E-03	Ν
rs10810657	9p22.2	BNC2	A/T	0.107	0.027	5.7E-05	-0.085	0.029	3.0E-03	Ν
rs191177147	3q28	LPP	G/T	0.097	0.03	1.0E-03	0.083	0.032	9.1E-03	Ν
rs9275642	6p21.32	HLA-DQA2	C/T	-0.212	0.096	2.7E-02	-2.812	1.406	4.6E-02	Ν
rs73183643	7q22.1	CUX1	G/A	-0.115	0.031	2.3E-04	-0.132	0.034	8.8E-05	Ν
rs11993814	8q21.13	ZBTB10	C/T	-0.06	0.03	4.5E-02	-0.083	0.032	9.5E-03	Ν
rs2776353	21q22.3	LINC00111	A/T	0.096	0.029	7.7E-04	-0.106	0.031	5.5E-04	Ν
rs10425559	19p13.3	PLIN3	G/A	-0.007	0.035	8.4E-01	0.004	0.035	9.0E-01	Ν
rs1050529	6p21.33	HLA-B	C/T	-0.118	0.107	2.7E-01	-0.312	0.179	8.2E-02	Ν
rs7776701	7p12.3	TNS3	C/T	-0.017	0.026	5.1E-01	-0.005	0.028	8.7E-01	Ν
rs9267650	6p21.3	NEU1	A/T	-0.086	0.066	2.0E-01	0.076	0.072	2.9E-01	Ν
rs7907606	10q24.3	OBFC1	T/G	-0.077	0.034	2.4E-02	0.073	0.037	4.9E-02	Ν
rs4710154	6q27	MIR3939	A/T	-0.058	0.028	4.3E-02	0.068	0.031	2.7E-02	Ν
rs2294214	6p22.3	CASC15	A/C	-0.12	0.03	5.9E-05	0.135	0.032	2.9E-05	N

Supplementary Table 14. Adjustment analysis for pigmentation phenotype in stage 2.

<sup>a</sup> Analysis was adjusted by hair color, tanning ability during adolescence, and number of blistering sunburns in the logistic regression association model. " $\beta$ " refers to effect size, "SE" to standard error, "Adj" to adjusted, "PC" to previously confirmed susceptibility locus, and "N" to novel susceptibility locus.

						Stage 2			
SNP	Region	Gene	Maj/ min	<i>P</i> for SNP	Int <i>P</i> hair color	Int <i>P</i> sunburn	Int P tan	Int <i>P</i> UVB	Note
rs12203592	6p25.3	IRF4	C/T	1.5E-11	0.0147	0.1269	0.0231	0.1574	PC
rs1805007	16q24.3	MC1R	C/T	1.9E-09	0.2617	0.0553	0.4527	0.8077	PC
rs12210050	6p25.3	EXOC2	C/T	4.5E-07	0.34	0.9371	0.2096	0.5712	PC
rs214785	20p13	TGM3	T/C	1.2E-05	0.6537	0.1576	0.7358	0.4297	PC
rs35407	5p13.2	SLC45A2	G/A	7.4E-05	0.7456	0.7548	0.9102	0.0587	PC
rs6059655	20q11.22	RALY	G/A	2.8E-03	0.0894	0.5644	0.5252	0.1697	PC
rs57142672	1p36.13	RCC2	A/G	3.4E-03	0.9476	0.6069	0.3238	0.5949	PC
rs73635312	10p14	GATA3	G/A	2.3E-04	0.5108	0.6284	0.8583	0.3658	PC
rs1126809	11q14.3	TYR	G/A	1.3E-01	0.5733	0.1951	0.0711	0.06	PC
rs421284	5p15.33	CLPTM1L	T/C	2.8E-02	0.1991	0.2146	0.4098	0.8742	PC
rs2080303	2q33.1	ALS2CR12	C/T	1.9E-04	0.9261	0.7925	0.0471	0.912	PC
rs61824911	1q42.13	RHOU	A/G	1.4E-01	0.5005	0.3275	0.8657	0.4982	PC
rs12916300	15q13.1	OCA2	T/C	5.0E-04	0.1479	0.1949	0.148	0.4302	PC
rs141115006	8q22.2	RGS22	C/T	5.1E-03	0.9034	0.6912	0.074	0.8917	PC
rs10093547	8q21.11	ZFHX4	T/G	2.7E-04	0.0971	0.811	0.859	0.4303	PC
rs7874604	9p21.3	CDKN2B	T/C	3.8E-03	0.1196	0.0281	0.4439	0.2258	PC
rs11170164	12q13.13	KRT5	C/T	3.7E-06	0.9566	0.1993	0.9099	0.6325	PC
rs78378222	17p13.1	TP53	T/G	4.0E-03	0.8911	0.7125	0.0881	0.1847	PC
rs157935	7q32.3	KLF14	T/G	2.3E-02	0.4197	0.7012	0.6323	0.5127	PC
rs57244888	2p24.3	MYCN	T/C	4.4E-01	0.8525	0.9742	0.5191	0.8522	PC
rs7335046	13q32.3	UBAC2	C/G	2.5E-04	0.4801	0.7122	0.8656	0.1632	PC
rs2116709	3p13	FOXP1	A/T	6.1E-04	0.017	0.8123	0.0234	0.0453	Ν
rs10810657	9p22.2	BNC2	A/T	5.7E-05	0.0268	0.6183	0.1503	0.4562	Ν
rs191177147	3q28	LPP	G/T	1.0E-03	0.0002	0.9832	0.7416	0.8174	Ν
rs9275642	6p21.32	HLA-DQA2	C/T	2.7E-02	0.8354	0.4512	0.2499	0.9685	Ν
rs73183643	7q22.1	CUX1	G/A	2.3E-04	0.7846	0.9846	0.7022	0.6498	Ν
rs11993814	8q21.13	ZBTB10	C/T	4.5E-02	0.4267	0.4773	0.6854	0.7465	Ν
rs2776353	21q22.3	LINC00111	A/T	7.7E-04	0.6476	0.7809	0.1028	0.2998	Ν
rs10425559	19p13.3	PLIN3	G/A	8.4E-01	0.0058	0.9691	0.9614	0.5868	Ν
rs1050529	6p21.33	HLA-B	C/T	2.7E-01	0.2365	0.162	0.7724	0.1615	Ν
rs7776701	7p12.3	TNS3	C/T	5.1E-01	0.6746	0.4589	0.3433	0.0521	Ν
rs9267650	6p21.3	NEU1	A/T	2.0E-01	0.7232	0.4631	0.688	0.7657	Ν
rs7907606	10q24.3	OBFC1	T/G	2.4E-02	0.0505	0.5912	0.0194	0.397	Ν
rs4710154	6q27	MIR3939	A/T	4.3E-02	0.0021	0.9838	0.8908	0.5559	Ν
rs2294214	6p22.3	CASC15	A/C	5.9E-05	0.2909	0.3244	0.3393	0.0895	Ν

Supplementary Table 15. Interaction analysis for hair color, number of sunburns, tanning ability, and UVB exposure in stage 2.

"P for SNP" refers to logistic regression P-value in stage 2, "Int" to interaction, "PC" to previously confirmed susceptibility locus, and "N" to novel susceptibility locus.

						Stage 1					:	Stage 2						Meta-a	inalysis		
SNP	Region	Gene	Maj/ min	MAF	Р	OR	95% CI	Avg r <sup>2</sup>	MAF	Р	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	P <sub>het</sub>	ľ
							(1.06,					(0.98,							(1.06,		
rs9420907	10q24.33	OBFC1	A/C	0.14	2.0×10 <sup>-7</sup>	1.10	1.14)	1.00	0.14	1.7×10 <sup>-1</sup>	1.05	1.13)	1.00	1.00	1.00	1.00	1.2×10 <sup>-7</sup>	1.09	1.13)	0.24	29%
rs9419958	10q24.33	OBFC1	C/T	0.14	1.9×10 <sup>-7</sup>	1.10	(1.06, 1.14) (1.06	1.00	0.14	1.7×10 <sup>-1</sup>	1.05	(0.98, 1.13) (1	1.00	1.00	1.00	1.00	1.1×10 <sup>-7</sup>	1.09	(1.06, 1.13) (1.06	0.23	30%
rs4387287	10q24.33	OBFC1	C/A	0.17	6.8×10 <sup>-8</sup>	1.10	(1.00, 1.14) (1.06	0.98	0.17	3.6×10 <sup>-2</sup>	1.07	(1, 1.15) (0.99	0.98	0.94	1.00	0.97	6.7×10 <sup>-9</sup>	1.09	(1.00, 1.13) (1.06	0.52	0%
rs10786775	10q24.33	OBFC1	C/G	0.10	4.4×10 <sup>-6</sup>	1.11	1.15)	1.00	0.10	6.9×10 <sup>-2</sup>	1.08	1.17)	0.99	1.00	1.00	1.00	7.0×10 <sup>-7</sup>	1.10	1.14)	0.62	0%

### Supplementary Table 16. Meta-analysis for 4 SNPs within OBFC1, in linkage disequilibrium with index SNP rs7907606

For each SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 1 minor allele frequency (MAF), *P* value for stage 1, odds ratio (for stage 1 overall, plus 95% CI), average imputation r<sup>2</sup> (for stage 1), MAF for stage 2, *P* value for stage 2, odds ratio (for stage 2 overall, plus 95% CI), average imputation r<sup>2</sup> (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), average imputation r<sup>2</sup> (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and *P*<sub>het</sub> and *I*<sup>2</sup> for meta-analysis. *P* values generated via logistic regression.

						Stage 1					9	Stage 2						Meta-a	inalysis		
SNP	Region	Gene	Maj/ min	MAF	Р	OR	95% CI	Avg r <sup>2</sup>	MAF	Ρ	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	P <sub>het</sub>	I <sup>2</sup>
							(0.89,					(0.88,							(0.9,		
rs1464510	3q28	LPP	C/A	0.44	7.3×10 <sup>-11</sup>	0.92	0.94)	1.00	0.45	1.8×10 <sup>-3</sup>	0.92	0.97)	0.98	0.97	0.99	0.98	5.5×10 <sup>-13</sup>	0.92	0.94)	0.83	0%
rs9860547	3q28	LPP	G/A	0.46	5.3×10 <sup>-10</sup>	1.09	(1.06, 1.12)	0.97	0.46	2.8×10 <sup>-3</sup>	1.08	(1.03 <i>,</i> 1.14)	0.92	0.96	0.97	0.95	5.5×10 <sup>-12</sup>	1.09	(1.06, 1.11)	0.85	0%

Supplementary Table 17. Meta-analysis for 2 SNPs within LPP, in linkage disequilibrium with index SNP rs191177147

For each *LPP* SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 1 minor allele frequency (MAF), *P* value for stage 1, odds ratio (for stage 1 overall, plus 95% CI), average imputation  $r^2$  (for stage 1, odds ratio 2, *P* value for stage 2, odds ratio (for stage 2 overall, plus 95% CI), average imputation  $r^2$  (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), average imputation  $r^2$  (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and  $P_{het}$  and  $I^2$  for meta-analysis. *P* values generated via logistic regression.

					St	tage 1						Stage 2						Meta-an	alysis		
SNP	Region	Gene	Maj/ min	MAF	Р	OR	95% CI	Avg r <sup>2</sup>	MAF	Р	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	P <sub>het</sub>	ľ
rs7255265	19p13.3	TICAM1	C/T	0.37	3.7×10 <sup>-7</sup>	0.93	(0.91 <i>,</i> 0.96)	0.99	0.35	8.6×10 <sup>-1</sup>	0.99	(0.94 <i>,</i> 1.06)	0.37	0.97	1.00	0.78	2.5×10 <sup>-6</sup>	0.94	(0.92 <i>,</i> 0.97)	0.06	72%

Supplementary Table 18. Meta-analysis for 1 SNP within TICAM1, in linkage disequilibrium with index SNP rs10425559

For each SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 1 minor allele frequency (MAF), *P* value for stage 1, odds ratio (for stage 1 overall, plus 95% CI), average imputation r<sup>2</sup> (for stage 1), MAF for stage 2, *P* value for stage 2, odds ratio (for stage 2 overall, plus 95% CI), average imputation r<sup>2</sup> (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and *P*<sub>het</sub> and *I*<sup>2</sup> for meta-analysis. *P* values generated via logistic regression.

Supplementary Table 19. Meta-analysis for 1	SNP in linkage disequ	uilibrium with index SNP rs4	1710154
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				Stage 1					Stage 2								Meta-analysis				
SNP	Region	Gene	Maj/ min	MAF	Р	OR	95% CI	Avg r <sup>2</sup>	MAF	Р	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	P <sub>het</sub>	ľ
rs9355610	6q27	RNASET2	G/A	0.34	2.6×10 <sup>-7</sup>	1.07	(1.05, 1.1)	1.00	0.34	4.6×10 <sup>-2</sup>	1.05	(1, 1.11)	1.00	0.98	1.00	0.99	3.7×10 <sup>-8</sup>	1.07	(1.04, 1.1)	0.56	0%

For each SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 1 minor allele frequency (MAF), *P* value for stage 1, odds ratio (for stage 1 overall, plus 95% CI), average imputation r<sup>2</sup> (for stage 1), MAF for stage 2, *P* value for stage 2, odds ratio (for stage 2 overall, plus 95% CI), average imputation r<sup>2</sup> (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and *P*<sub>het</sub> and *I*<sup>2</sup> for meta-analysis. *P* values generated via logistic regression.

				Stage 1							:	Stage 2	Meta-analysis								
SNP	Region	Gene	Maj/ min	MAF	Р	OR	95% CI	Avg r <sup>2</sup>	MAF	Ρ	OR	95% CI	Affy	Illum	Omni	Avg r <sup>2</sup>	Р	OR	95% CI	P <sub>het</sub>	I <sup>2</sup>
							(0.87.					(0.86.							(0.88.		
rs62543565	9p22.2	BNC2	A/C	0.41	1.1×10 <sup>-13</sup>	0.90	0.93)	0.90	0.40	3.5×10 <sup>-4</sup>	0.91	0.96)	0.89	0.87	0.88	0.88	2.2×10 <sup>-16</sup>	0.90	0.92)	0.84	0.0%
rs10756819	9p22.2	BNC2	A/G	0.34	1.5×10 <sup>-5</sup>	0.94	0.97)	1.00	0.34	7.7×10 <sup>-3</sup>	0.93	0.98)	0.89	1.00	1.00	0.96	4.2×10 <sup>-7</sup>	0.94	0.96)	0.67	0.0%
rs2153271	9p22.2	BNC2	T/C	0.42	1.5×10 <sup>-11</sup>	0.91	0.94)	1.00	0.42	4.8×10 <sup>-5</sup>	0.90	0.95)	0.98	1.00	1.00	0.99	4.3×10 <sup>-15</sup>	0.91	0.93)	0.60	0%
rs12350739	9p22.2	BNC2	A/G	0.44	2.3×10 <sup>-12</sup>	0.91	0.93)	0.97	0.43	7.2×10 <sup>-6</sup>	0.89	0.93)	0.90	0.97	0.96	0.94	1.4×10 <sup>-16</sup>	0.90	0.93)	0.41	0.0%

## Supplementary Table 20. Meta-analysis for 4 SNPs within BNC2, in linkage disequilibrium with index SNP rs10810657

For each *BNC2* SNP, we report rsID, genetic locus, gene, major and minor alleles, stage 1 minor allele frequency (MAF), *P* value for stage 1, odds ratio (for stage 1 overall, plus 95% CI), average imputation  $r^2$  (for stage 1, MAF for stage 2, *P* value for stage 2, odds ratio (for stage 2 overall, plus 95% CI), average imputation  $r^2$  (for Affy, Illumina, Omni, stage 2 overall), *P* value for meta-analysis, odds ratio (for meta-analysis overall, plus 95% CI), and *P*<sub>het</sub> and *I*<sup>2</sup> for meta-analysis. *P* values generated via logistic regression.