

Supporting information

S1 table: Summary of candidate genes in the study associated with leprosy

S2 table: Ethnicity distribution of the study groups

S3 table: Analysis of association between SNPs and leprosy^a

S4 table: Genotypic distributions of the study groups

S5 table: Analysis of association between SNPs and leprosy onset early cases

S6 table: Analysis of association between SNPs and leprosy WHO classifications

S7 table: Analysis of association between SNPs and leprosy subtypes

S8 table: Analysis of association between SNPs and reversal reaction in BB, BT and

BL leprosy cases

S9 table: Analysis of association between SNPs and erythema nodosum leprosum in LL

and BL leprosy cases

S10 table: Analysis of association between SNPs and leprosy reaction states

Supplementary Table 1. Summary of candidate genes in the study associated with leprosy

SNP	Gene	Functional Consequence	MAF ^a	Molecular function for genes
rs76418789	<i>IL23R</i>	missense	0.011	IL23R was critical for optimal levels of pattern-recognition receptor-induced signaling and cytokines in human monocyte-derived macrophages ¹
rs146466242	<i>FLG</i>	stop gained	0.004	Encoded filaggrin, filaggrin plays a crucial role in the barrier function of epidermis ²
rs2221593	<i>BATF3</i>	upstream	0.161	Plays a central role in the development of conventional dendritic cells, which in turn are critical for optimal priming of CD8+ T cells ³
rs2058660	<i>IL18RAP/IL18RI</i>	intron	0.694	Plays an important role in IFN- γ production ⁴
rs73058713	<i>CDH18</i>	intron	0.178	Involves in neuronal development and govern metabolic processes in later life ^[5]

rs2275606	<i>RAB32</i>	ncRNA intronic	0.134	Key regulator of a host defense pathway that can restrict intracellular bacterial pathogens ⁶
rs4720118	<i>BBS9</i>	intron	0.672	Involves in parathyroid hormone action in bones ⁷
rs55894533	<i>CTSB</i>	intergenic	0.409	Associated with the migration ability of keratinocytes and unobstructed migration of keratinocytes ⁷
rs6478108	<i>TNFSF15</i>	intron	0.715	Involved in mediating the switch from Th1 cells to Th2 cells ⁸
rs10817758	<i>I-Dec</i>	intron	0.182	Acts as an accelerator in tumor progression ^[9]
rs780668	<i>SLC29A3</i>	missense	0.480	Encodes ENT3, a key role in maintaining T cell homeostasis by supporting the proliferation and survival of T cells ¹⁰
rs663743	<i>CCDC88</i> <i>B</i>	5 prime UTR variant	0.198	A key role for mobility and inflammatory functions of dendritic cells ¹¹
rs3764147	<i>LACCI</i>	missense	0.306	Associated with autoimmunity and granulomatous disease diseases ¹²

rs142179458	<i>HIF1A</i>	missense	0.005	Plays a role in immune reactions and chaperone-mediated autophagy ¹³
rs9302752	<i>NOD2</i>	upstream	0.500	pro-inflammatory ¹⁴
rs149308743	<i>CARD9</i>	missense	<0.001	The adaptor CARD9 functions downstream of C-type lectin receptors for the sensing of microbial infection ¹⁵
rs145562243	<i>NCKIPSD</i>	missense	<0.0001	involved in the Rho signaling pathway ¹⁶

^a Minor allele frequency (MAF) from the 1000 Genomes Project¹⁷, variants with MAF <0.05 were regarded as rare.

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Supplementary table 2: Ethnicity distribution of the study groups

Ethnicity	Leprosy (n=1344)	Heredit- related Contacts (n=1908)	First-degree family members (n=1560)	Non-heredit- related Contacts (n=824)
Han	711(52.9)	1106(58.0)	872(55.9)	453(55.0)
Miao	169(12.6)	229(12.0)	197(12.6)	106(12.9)
Yi	144(10.7)	169(8.9)	141(9.0)	82(10.0)
Buyi	83(6.2)	52(2.7)	52(3.3)	26(3.2)
Tujia	35(2.6)	64(3.4)	42(2.7)	25(3.0)
Zhuang	34(2.5)	73(3.8)	61(3.9)	28(3.4)
Dai	39(2.9)	60(3.1)	54(3.5)	36(4.4)
Others	129(9.6)	155(8.1)	141(9.0)	68(8.3)

Supplementary table 3: Analysis of association between SNPs and leprosy^a

SNP	Gene	Minor allele	F_A	F_U ^b	F_U ^c	F_U ^d	F_U ^e	OR (95% CI) ^b	P ^b	OR (95% CI) ^c	P ^c	OR (95% CI) ^d	P ^d	OR (95% CI) ^e	P ^e
rs2275606	<i>RAB32</i>	A	0.29	0.28	0.28	0.28	0.27	1.08(0.98 -1.20)	0.12	1.07 (0.96- 1.19)	0.24	1.05 (0.94- 1.18)	0.37	1.12 (0.98-1.29)	0.10
rs142179458	<i>HIF1A</i>	A	0.05	0.05	0.05	0.05	0.05	1.02(0.82 -1.26)	0.88	1.00 (0.80- 1.26)	0.98	0.97 (0.77- 1.24)	0.82	1.05 (0.79-1.41)	0.74
rs3764147	<i>LACCI</i>	G	0.41	0.34	0.36	0.37	0.30	1.362(1.24 -1.50)	1.62×10 ⁻¹⁰	1.26(1.14- 1.40)	<0.0001	1.22(1.10- 1.36)	0.0002	1.26 (1.14-1.40)	<0.0001
rs55894533	<i>CTSB</i>	C	0.47	0.45	0.45	0.45	0.45	1.08(0.99 -1.19)	0.09	1.08 (0.98- 1.19)	0.13	1.08 (0.97- 1.20)	0.16	1.10 (0.97-1.24)	0.16
rs6478108	<i>TNFSF15</i>	T	0.40	0.43	0.43	0.42	0.44	0.89(0.81 -0.97)	0.01	0.90 (0.81- 0.99)	0.03	0.92 (0.83- 1.03)	0.13	0.87 (0.76-0.98)	0.02
rs73058713	<i>CDH18</i>	A	0.14	0.13	0.13	0.13	0.13	1.06(0.93 -1.22)	0.36	1.06 (0.92-1.22)	0.44	1.09 (0.94-1.27)	0.24	1.08 (0.90-1.29)	0.40
rs780668	<i>SLC29A3</i>	A	0.50	0.45	0.46	0.46	0.44	1.20(1.10 -1.32)	9.48×10 ⁻⁵	1.17 (1.06-1.29)	0.002	1.15(1.04-1.28)	0.007	1.28 (1.13-1.45)	<0.0001
rs10817758	<i>DECI</i>	T	0.44	0.46	0.46	0.46	0.45	0.94 (0.86 -1.04)	0.22	0.93 (0.84-1.02)	0.13	0.92 (0.83-1.02)	0.11	0.98 (0.87-1.11)	0.79
rs146466242	<i>FLG</i>	A	0.02	0.02	0.02	0.02	0.02	1.00(0.71 -1.40)	0.98	1.00 (0.69-1.44)	1.00	0.97 (0.66-1.41)	0.86	0.99 (0.63-1.55)	0.96
rs9302752	<i>NOD2</i>	C	0.27	0.24	0.25	0.25	0.22	1.18(1.06 -1.31)	0.002	1.11 (0.99-1.24)	0.07	1.12 (1.00-1.26)	0.06	1.35 (1.17-1.56)	<0.0001
rs2058660	<i>IL18RAP/IL18R1</i>	C	0.42	0.43	0.42	0.42	0.43	0.96 (0.88 -1.06)	0.45	0.98 (0.89-1.08)	0.69	1.00 (0.90-1.11)	1.00	0.93 (0.82-1.05)	0.25

rs4720118	<i>BBS9</i>	T	0.30	0.29	0.29	0.29	0.30	1.06(0.96 -1.17)	0.27	1.07 (0.96-1.19)	0.23	1.09 (0.97-1.22)	0.14	1.04 (0.91-1.18)	0.61
rs663743	<i>CCDC88B</i>	A	0.23	0.21	0.21	0.22	0.21	1.13(1.01 -1.26)	0.03	1.12 (0.99-1.26)	0.06	1.10 (0.97-1.24)	0.14	1.15 (0.99-1.34)	0.06
rs76418789	<i>IL23R</i>	A	0.05	0.04	0.04	0.04	0.03	1.14(0.91 -1.44)	0.24	1.07 (0.84-1.36)	0.57	1.04 (0.81-1.34)	0.74	1.35 (0.98-1.87)	0.07

Abbreviations: SNP, single nucleotide polymorphism; F_A, minor allele frequency in patients; F_U, minor allele frequency in control subjects; OR, odds ratio with respect to the minor allele; CI, confidence interval.

^a Association tested with χ^2 test or Fisher test.

^b All HHCs as control subjects.

^c Heredity-related Contacts as control subjects.

^d First - degree family members as control subjects.

^e non-Heredity-related Contacts as control subjects.

Supplementary table 4: Genotypic distributions of the study groups

SNP	Gene	Mino r allele	aa/Aa/AA ^a	Cases (n=1344)	Controls ^b (n=2732)	Controls ^c (n=1908)	Controls ^d (n=1560)	Controls ^e (n=824)
rs2275606	<i>RAB32</i>	A	121/545/6 78	206/1098/ 1428	150/766/ 992	127/626/8 07	56/332/43 6	
rs1421794 58	<i>HIF1A</i>	A	4/122/121 8	10/240/24 82	8/168/17 32	7/141/141 2	2/72/750	
rs3764147	<i>LACC1</i>	G	225/665/4 54	315/1240/ 1177	249/875/ 784	208/730/6 22	66/365/39 3	
rs5589453 3	<i>CTSB</i>	C	300/663/3 81	542/1374/ 816	382/956/ 570	313/782/4 65	160/418/2 46	
rs6478108	<i>TNFSF15</i>	T	238/609/4 97	527/1311/ 894	367/908/ 633	299/722/5 39	160/403/2 61	
rs7305871 3	<i>CDH18</i>	A	24/332/98 8	59/614/20 59	38/438/1 432	26/356/11 78	21/176/62 7	
rs780668	<i>SLC29A3</i>	A	339/655/3 50	522/1415/ 795	380/982/ 546	316/805/4 39	142/433/2 49	
rs1081775 8	<i>DECI</i>	T	252/682/4 10	563/1364/ 805	400/956/ 552	325/792/4 43	163/408/2 53	
rs1464662 42	<i>FLG</i>	A	3/44/1297	3/96/2633	2/67/183 9	2/56/1502	1/29/794	
rs9302752	<i>NOD2</i>	C	103/524/7 17	154/1007/ 1571	119/721/ 1068	102/575/8 83	35/286/50 3	
rs2058660	<i>IL18RAP/IL18 RI</i>	C	225/670/4 49	523/1279/ 930	361/887/ 660	287/726/5 47	162/392/2 70	
rs4720118	<i>BBS9</i>	T	124/571/6 49	234/1132/ 1366	154/802/ 952	124/647/7 89	80/330/41 4	
rs663743	<i>CCDC88B</i>	A	70/482/79 2	140/870/1 722	94/621/1 193	80/511/96 9	46/249/52 9	
rs7641878 9	<i>IL23R</i>	A	5/112/122 7	6/206/252 0	4/154/17 50	3/130/142 7	2/52/770	

Abbreviations: SNP, single nucleotide polymorphism;

^a aa/Aa/AA, the numbers of cases and controls with minor allele homozygote/heterozygote/major allele homozygote genotypes, respectively.

^b All HHCs as control subjects.

^c Heredity-related Contacts as control subjects.

^d First - degree family members as control subjects.

^e non-Heredity-related contacts as control subjects.

Supplementary table 5: Analysis of association between SNPs and leprosy onset early cases

SNP	Gene	Minor allele	aa/Aa/AA ^a	F_A	F_U	OR (95% CI) ^b	P ^b	Risk allele	OR (95% CI) ^c	P ^c	
			Cases (n=44)	Controls (n=1300)							
rs2275606	<i>RAB32</i>	A	6/19/19	115/526/659	0.35	0.29	1.33 (0.85-2.07)	0.21	A	1.27 (0.81-1.97)	0.30
rs142179458	<i>HIF1A</i>	A	1/8/35	3/114/1183	0.11	0.05	2.65 (1.34-5.25)	0.004	A	2.31 (1.16-4.62)	0.02
rs3764147	<i>LACCI</i>	G	5/22/17	220/643/437	0.36	0.42	0.80 (0.51-1.24)	0.32	A	1.26 (0.81-1.98)	0.31
rs55894533	<i>CTSB</i>	C	8/23/13	292/640/368	0.44	0.47	0.89 (0.58-1.37)	0.61	A	1.13 (0.74-1.74)	0.56
rs6478108	<i>TNFSF15</i>	T	7/25/12	231/584/485	0.44	0.40	1.18 (0.77-1.81)	0.44	T	1.16 (0.76-1.75)	0.49
rs73058713	<i>CDH18</i>	A	0/13/31	24/319/957	0.15	0.14	1.06 (0.58-1.92)	0.86	A	1.09 (0.59-2.00)	0.79
rs780668	<i>SLC29A3</i>	A	14/21/9	325/634/341	0.56	0.49	1.29 (0.84-1.98)	0.25	A	1.26 (0.83-1.92)	0.28
rs10817758	<i>DECI</i>	T	10/25/9	242/657/401	0.51	0.44	1.34 (0.87-2.05)	0.18	T	1.32 (0.85-2.04)	0.22
rs146466242	<i>FLG</i>	A	0/2/42	3/42/1255	0.02	0.02	1.24 (0.30-5.17)	0.77	A	1.27 (0.33-4.97)	0.73
rs9302752	<i>NOD2</i>	C	5/20/19	98/504/698	0.34	0.27	1.40 (0.90-2.20)	0.14	C	1.48 (0.95-2.33)	0.09
rs2058660	<i>IL18RAP/IL18R1</i>	C	10/19/15	215/651/434	0.44	0.42	1.12 (0.73-1.72)	0.61	C	1.14 (0.74-1.77)	0.55
rs4720118	<i>BBS9</i>	T	4/16/24	120/555/625	0.27	0.31	0.85 (0.53-1.37)	0.51	C	1.16 (0.72-1.88)	0.55
rs663743	<i>CCDC88B</i>	A	4/13/27	66/469/765	0.24	0.23	1.04 (0.63-1.72)	0.87	A	1.00 (0.61-1.65)	0.99
rs76418789	<i>IL23R</i>	A	0/2/42	5/110/1185	0.02	0.05	0.48 (0.12-1.98)	0.30	G	2.01 (0.49-8.27)	0.33

Abbreviations: SNP, single nucleotide polymorphism; F_A, minor allele frequency in patients diagnosed at the age<14-year; F_U, minor allele frequency in patients diagnosed at the age≥14-year; OR, odds ratio with respect to the minor allele; CI,

confidence interval.

^a aa/Aa/AA, the numbers of cases and controls with minor allele homozygote/heterozygote/major allele homozygote genotypes, respectively

^b Association tested with χ^2 test or Fisher test.

^c Association tested with logistic regression model adjusted for sex and ethnicity.

Supplementary table 6: Analysis of association between SNPs and leprosy WHO classifications

SNP	Gene	Minor allele	aa/Aa/AA ^a	F_A	F_U	OR (95% CI) ^b	P ^b	Risk allele	OR (95% CI) ^c	P ^c	
			Cases (n=173)	Controls (n=1171)							
rs2275606	<i>RAB32</i>	A	12/73/88	109/472/590	0.28	0.29	0.93 (0.73-1.199)	0.59	G	1.07 (0.83-1.37)	0.62
rs142179458	<i>HIF1A</i>	A	0/18/155	4/104/1063	0.05	0.05	1.09 (0.66-1.82)	0.73	A	1.13 (0.68-1.88)	0.64
rs3764147	<i>LACCI</i>	G	32/78/63	193/587/391	0.41	0.42	0.98 (0.78-1.23)	0.86	A	1.02 (0.81-1.28)	0.89
rs55894533	<i>CTSB</i>	C	39/86/48	261/577/333	0.47	0.47	1.02 (0.81-1.28)	0.87	C	1.02 (0.81-1.28)	0.88
rs6478108	<i>TNFSF15</i>	T	27/90/56	211/519/441	0.42	0.40	1.06 (0.84-1.34)	0.61	T	1.03 (0.83-1.29)	0.77
rs73058713	<i>CDH18</i>	A	2/48/123	22/284/865	0.15	0.14	1.09 (0.79-1.49)	0.61	A	1.10 (0.79-1.52)	0.57
rs780668	<i>SLC29A3</i>	A	41/88/44	298/567/306	0.49	0.50	0.98 (0.78-1.23)	0.86	A	1.01 (0.81-1.26)	0.93
rs10817758	<i>DECI</i>	T	38/85/50	214/597/360	0.47	0.44	1.12 (0.89-1.40)	0.33	T	1.14 (0.90-1.44)	0.27
rs146466242	<i>FLG</i>	A	0/6/167	3/38/1130	0.02	0.02	0.92 (0.39-2.18)	0.85	A	1.06 (0.46-2.43)	0.89
rs9302752	<i>NOD2</i>	C	8/71/94	95/453/623	0.25	0.27	0.89 (0.68-1.15)	0.37	T	1.08 (0.83-1.40)	0.57
rs2058660	<i>IL18RAP/IL18R1</i>	C	36/82/55	189/588/394	0.45	0.41	1.14 (0.91-1.43)	0.25	C	1.15 (0.91-1.45)	0.24
rs4720118	<i>BBS9</i>	T	17/84/72	107/487/577	0.34	0.30	1.21 (0.95-1.54)	0.12	T	1.21 (0.95-1.54)	0.13
rs663743	<i>CCDC88B</i>	A	9/70/94	61/412/698	0.25	0.23	1.16 (0.90-1.50)	0.28	A	1.15 (0.88-1.50)	0.29
rs76418789	<i>IL23R</i>	A	0/10/163	5/102/1064	0.03	0.05	0.59 (0.31-1.14)	0.11	G	1.62 (0.84-3.11)	0.15

Abbreviations: SNP, single nucleotide polymorphism; F_A, minor allele frequency in paucibacillary leprosy cases; F_U, minor allele frequency in multibacillary leprosy cases; OR, odds ratio with respect to the minor allele; CI, confidence interval.

^a aa/Aa/AA, the numbers of cases and controls with minor allele

homozygote/heterozygote/major allele homozygote genotypes, respectively.

^b Association tested with χ^2 test or Fisher test.

^c Association tested with logistic regression model adjusted for sex, ethnicity and age at diagnosis.

Supplementary table 7: Analysis of association between SNPs and leprosy

subtypes

SNP	Gene	Minor allele	aa/Aa/AA ^a		F_A	F_U	OR (95% CI) ^b	P ^b	Risk allele	OR (95% CI) ^c	P ^c
			Cases (n=391)	Controls (n=841)							
rs7641878 9	<i>IL23R</i>	A	0/28/363	5/75/761	0.04	0.05	0.70(0.45-1.08)	0.10	G	1.37(0.89-2.11)	0.15
rs1464662 42	<i>FLG</i>	A	1/11/379	2/26/813	0.02	0.02	0.93(0.48-1.80)	0.83	A	1.06(0.57-1.98)	0.86
rs2058660	<i>IL18RAP/IL18RI</i>	C	79/184/128	132/426/28 3	0.44	0.41	1.12(0.94-1.33)	0.20	C	1.13(0.94-1.34)	0.19
rs7305871 3	<i>CDH18</i>	A	7/94/290	13/206/622	0.14	0.14	1.00(0.78-1.28)	0.99	A	1.01(0.78-1.29)	0.97
rs2275606	<i>RAB32</i>	A	25/166/200	82/333/426	0.28	0.30	0.91 (0.75-1.10)	0.33	G	1.09 (0.90-1.31)	0.39
rs4720118	<i>BBS9</i>	T	36/175/180	77/342/422	0.32	0.29	1.10(0.92-1.33)	0.29	T	1.10(0.92-1.32)	0.31
rs5589453 3	<i>CTSB</i>	C	77/216/98	199/395/24 7	0.47	0.47	1.01(0.85-1.19)	0.94	C	1.01(0.85-1.20)	0.91
rs6478108	<i>TNFSF15</i>	T	71/187/133	151/361/32 9	0.42	0.39	1.12(0.94-1.33)	0.21	T	1.09(0.92-1.29)	0.32
rs1081775 8	<i>I-Dec</i>	T	79/182/130	163/436/24 2	0.43	0.45	0.93(0.78-1.10)	0.40	C	1.07(0.90-1.27)	0.46
rs780668	<i>SLC29A3</i>	A	91/192/108	222/400/21 9	0.48	0.50	0.91(0.77-1.08)	0.28	G	1.07(0.91-1.27)	0.42
rs663743	<i>CCDC88B</i>	A	19/144/228	43/300/498	0.23	0.23	1.02(0.83-1.25)	0.86	A	1.01(0.82-1.24)	0.92

rs3764147	<i>LACCI</i>	G	51/174/166	161/435/24 5	0.35	0.45	0.67(0.56- 0.79)	<0.0001	A	1.51(1.26- 1.81)	<0.0001
rs1421794 58	<i>HIF1A</i>	A	3/40/348	1/71/769	0.06	0.04	1.38(0.94- 2.01)	0.10	A	1.46(0.99- 2.14)	0.05
rs9302752	<i>NOD2</i>	C	21/151/219	75/330/436	0.25	0.29	0.82(0.68- 1.00)	<0.05	T	1.18(0.97- 1.44)	0.09

Abbreviations: SNP, single nucleotide polymorphism; F_A, minor allele frequency in tuberculoid cases (BT/TT); F_U, minor allele frequency in lepromatous cases (LL/BL); OR, odds ratio with respect to the minor allele; CI, confidence interval.

^a aa/Aa/AA, the numbers of cases and controls with minor allele homozygote/heterozygote/major allele homozygote genotypes, respectively.

^b Association tested with χ^2 test or Fisher test.

^c Association tested with logistic regression model adjusted for sex, ethnicity and age at diagnosis.

Supplementary table 8: Analysis of association between SNPs and reversal

reaction in BB, BT and BL leprosy cases

SNP	Gene	Minor allele	aa/Aa/AA ^a		F_A	F_U	OR (95% CI) ^b	P ^b	Risk allele	OR (95% CI) ^c	P ^c
			Cases (n=54)	Controls (n=696)							
rs76418789	<i>IL23R</i>	A	1/3/50	2/62/632	0.05	0.05	0.98 (0.38-2.47)	0.96	G	1.02 (0.40-2.56)	0.97
rs146466242	<i>FLG</i>	A	0/1/53	2/24/670	0.009	0.02	0.46 (0.06-3.38)	0.43	T	1.95 (0.28-13.67)	0.50
rs2058660	<i>IL18RAP/IL18R1</i>	C	8/30/16	114/344/238	0.43	0.41	1.06 (0.72-1.58)	0.76	C	1.07 (0.71-1.60)	0.75
rs73058713	<i>CDH18</i>	A	3/17/34	13/167/516	0.21	0.14	1.68 (1.04-2.73)	0.03	A	1.69 (1.04-2.76)	0.03
rs2275606	<i>RAB32</i>	A	5/21/28	69/275/352	0.29	0.30	0.95 (0.62-1.47)	0.83	G	1.06 (0.69-1.62)	0.78
rs4720118	<i>BBS9</i>	T	6/20/28	61/297/338	0.30	0.30	0.98 (0.64-1.50)	0.92	C	1.03 (0.67-1.58)	0.91
rs55894533	<i>CTSB</i>	C	7/28/19	159/347/190	0.39	0.48	0.70 (0.47-1.04)	0.07	A	1.46 (0.97-2.18)	0.07
rs6478108	<i>TNFSF15</i>	T	13/22/19	117/314/265	0.44	0.39	1.23 (0.83-1.83)	0.30	T	1.21 (0.83-1.77)	0.33
rs10817758	<i>I-Dec</i>	T	7/31/16	135/335/226	0.42	0.43	0.93 (0.62-1.38)	0.72	C	1.09 (0.73-1.62)	0.69
rs780668	<i>SLC29A3</i>	A	13/29/12	176/331/189	0.51	0.49	1.08 (0.73-1.59)	0.71	A	1.07 (0.73-1.58)	0.72
rs663743	<i>CCDC88B</i>	A	3/17/34	39/248/409	0.21	0.23	0.88 (0.55-1.43)	0.61	G	1.13 (0.70-1.82)	0.61
rs3764147	<i>LACCI</i>	G	6/18/30	112/347/237	0.28	0.41	0.55	0.007	A	1.85	0.007

rs142179458	<i>HIF1A</i>	A	0/7/47	3/71/622	0.06	0.06	(0.36-0.85)	0.68	A	(1.19-2.87)	0.73
							1.18			1.15	
							(0.53-2.63)			(0.52-2.55)	
rs9302752	<i>NOD2</i>	C	5/17/32	46/257/393	0.25	0.25	1.00	0.99	C	1.04	0.87
							(0.63-1.57)			(0.66-1.63)	

Abbreviations: SNP, single nucleotide polymorphism; F_A, minor allele frequency in reversal reaction in BB, BT and BL leprosy cases; F_U, minor allele frequency in BB, BT and BL leprosy cases free of reversal reaction; OR, odds ratio with respect to the minor allele; CI, confidence interval.

^a aa/Aa/AA, the numbers of cases and controls with minor allele homozygote/heterozygote/major allele homozygote genotypes, respectively.

^b Association tested with χ^2 test or Fisher test.

^c Association tested with logistic regression model adjusted for sex, ethnicity and age at diagnosis.

Supplementary table 9: Analysis of association between SNPs and erythema

nodosum leprosum in LL and BL leprosy cases

SNP	Gene	Minor allele	aa/Aa/AA ^a		F_A	F_U	OR (95% CI) ^b	P ^b	Risk allele	OR (95% CI) ^c	P ^c
			Cases (n=75)	Controls (n=713)							
rs76418789	<i>IL23R</i>	A	0/9/66	4/62/647	0.06	0.05	1.20 (0.60-2.53)	0.56	A	1.23 (0.61-2.47)	0.57
rs146466242	<i>FLG</i>	A	0/4/71	2/21/690	0.03	0.02	1.54 (0.53-4.47)	0.43	A	1.49 (0.54-4.11)	0.44
rs2058660	<i>IL18RAP/IL18R1</i>	C	17/35/23	104/366/243	0.46	0.40	1.26 (0.90-1.77)	0.17	C	1.26 (0.89-1.79)	0.19
rs73058713	<i>CDH18</i>	A	0/22/53	12/171/530	0.15	0.14	1.09 (0.67-1.75)	0.74	A	1.08 (0.66-1.77)	0.75
rs2275606	<i>RAB32</i>	A	6/28/41	73/282/358	0.27	0.30	0.85 (0.58-1.24)	0.39	G	1.16 (0.80-1.68)	0.44
rs4720118	<i>BBS9</i>	T	7/30/38	63/295/355	0.29	0.30	0.99 (0.68-1.43)	0.96	C	1.02 (0.71-1.48)	0.91
rs55894533	<i>CTSB</i>	C	16/33/26	167/342/204	0.43	0.47	0.85 (0.60-1.19)	0.34	A	1.18 (0.84-1.65)	0.33
rs6478108	<i>TNFSF15</i>	T	19/31/25	122/302/289	0.46	0.38	1.37 (0.98-1.93)	0.07	T	1.31 (0.95-1.81)	0.10
rs10817758	<i>I-Dec</i>	T	13/44/18	143/359/211	0.47	0.45	1.06 (0.76-1.49)	0.74	T	1.09 (0.77-1.54)	0.64
rs780668	<i>SLC29A3</i>	G	15/38/22	190/338/185	0.45	0.50	0.82 (0.58-1.15)	0.24	A	1.23 (0.88-1.72)	0.23
rs663743	<i>CCDC88B</i>	A	5/24/46	36/258/419	0.23	0.23	0.97 (0.65-1.45)	0.90	G	1.01 (0.67-1.52)	0.96
rs3764147	<i>LACCI</i>	G	21/38/16	134/369/210	0.53	0.45	1.42	0.04	G	1.44	0.04

rs142179458	<i>HIF1A</i>	A	1/5/69	0/59/654	0.05	0.04	(1.01-1.98)	0.76	A	(1.02-2.04)	0.67
							1.13			1.20	
							(0.51-2.53)			(0.53-2.71)	
rs9302752	<i>NOD2</i>	C	6/33/36	62/280/371	0.30	0.28	1.08	0.67	C	1.10	0.60
							(0.75-1.57)			(0.76-1.59)	

Abbreviations: SNP, single nucleotide polymorphism; F_A, minor allele frequency in erythema nodosum leprosum in LL and BL leprosy cases; F_U, minor allele frequency in LL and BL leprosy cases free of erythema nodosum leprosum; OR, odds ratio with respect to the minor allele; CI, confidence interval.

^a aa/Aa/AA, the numbers of cases and controls with minor allele homozygote/heterozygote/major allele homozygote genotypes, respectively.

^b Association tested with χ^2 test or Fisher test.

^c Association tested with logistic regression model adjusted for sex, ethnicity and age at diagnosis.

Supplementary table 10: Analysis of association between SNPs and leprosy

reaction states

SNP	Gene	Minor allele	aa/Aa/AA ^a		F_A	F_U	OR (95% CI) ^b	P ^b	Risk allele	OR (95% CI) ^c	P ^c
			Cases (n=79)	Controls (n=67)							
rs7641 8789	<i>IL23R</i>	A	0/9/70	1/4/62	0.06	0.04	1.29 (0.45- 3.718)	0.64	A	1.26 (0.45- 3.52)	0.66
rs1464 66242	<i>FLG</i>	A	0/4/75	0/1/66	0.03	0.007	3.46 (0.38- 31.29)	0.24	A	3.18 (0.34- 29.58)	0.31
rs2058 660	<i>IL18R</i> <i>AP/IL1</i>	C	17/39/ 23	11/36/20	0.46	0.43	1.13 (0.71- 1.79)	0.62	C	1.12 (0.69- 1.80)	0.65
rs7305 8713	<i>CDHI</i> 8	A	0/23/5 6	3/22/42	0.15	0.21	0.65 (0.35- 1.18)	0.16	C	1.69 (0.89- 3.23)	0.11
rs2275 606	<i>RAB32</i>	A	6/29/4 4	7/27/33	0.26	0.31	0.79 (0.48- 1.33)	0.38	G	1.20 (0.70- 2.04)	0.51
rs4720 118	<i>BBS9</i>	T	8/33/3 8	7/26/34	0.31	0.30	1.06 (0.64- 1.74)	0.83	T	1.07 (0.65- 1.75)	0.79
rs5589 4533	<i>CTSB</i>	C	17/35/ 27	10/34/23	0.44	0.40	1.15 (0.72- 1.83)	0.56	C	1.12 (0.70- 1.78)	0.64
rs6478 108	<i>TNFSF</i> 15	T	19/34/ 26	15/32/20	0.46	0.46	0.97 (0.61- 1.54)	0.90	C	1.02 (0.64- 1.62)	0.93
rs1081 7758	<i>I-Dec</i>	T	13/47/ 19	8/39/20	0.46	0.41	1.23 (0.77- 1.96)	0.38	T	1.31 (0.77- 2.21)	0.32
rs7806 68	<i>SLC29</i> <i>A3</i>	G	15/41/ 23	15/33/19	0.45	0.47	0.92 (0.58- 1.46)	0.72	A	1.12 (0.70- 1.80)	0.63
rs6637 43	<i>CCDC</i> <i>88B</i>	A	5/26/4 8	3/20/44	0.23	0.19	1.23 (0.70- 2.16)	0.48	A	1.34 (0.74- 2.40)	0.33
rs3764	<i>LACCI</i>	G	23/39/ 8	7/26/34	0.54	0.30	2.74	<0.00	G	2.75	<

147			17				(1.69- 4.44)	01		(1.65- 4.59)	0.0001
rs1421 79458	<i>HIF1A</i>	A	1/6/72	0/8/59	0.05	0.06	0.84 (0.31- 2.30)	0.73	G	1.15 (0.43- 3.05)	0.78
rs9302 752	<i>NOD2</i>	C	7/36/3 6	6/24/37	0.32	0.27	1.26 (0.76- 2.10)	0.37	C	1.32 (0.77- 2.25)	0.31

Abbreviations: SNP, single nucleotide polymorphism; F_A, minor allele frequency in erythema nodosum leprosum cases; F_U, minor allele frequency in reversal reaction cases; OR, odds ratio with respect to the minor allele; CI, confidence interval.

^a aa/Aa/AA, the numbers of cases and controls with minor allele

homozygote/heterozygote/major allele homozygote genotypes, respectively.

^b Association tested with χ^2 test or Fisher test.

^c Association tested with logistic regression model adjusted for sex, ethnicity and age at diagnosis.