# Multi-ancestry genome-wide association study identifies new asthma susceptibility loci that co-localize with immune cell enhancer histone marks 

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Supplementary Figures

## Supplementary Figure 1. Quantile-quantile (QQ) plots for the ancestry-specific and multi-ancestry meta-

 analyses of asthma. QQ plots of $P_{\text {random }}$ or $P_{\text {fixed }}$ values for all SNPs passing QC in a least two-thirds of studies.

Supplementary Figure 2. Manhattan plots for each meta-analysis in African ancestry, Japanese and Latino populations.
The -log10 Pfixed values for all SNPs passing QC in at least two-thirds of studies have been plotted; the dashed horizontal line corresponds to a $P$-value threshold of $10^{-5}$


Latino


## Supplementary Figure 3. Forest Plots of the lead SNPs at the 18 genome-wide significant loci

(nine loci harbouring new signals for asthma followed by nine known asthma loci, as shown in Table 1)

Odds-ratio (OR) and 95\% Confidence Intervals (CI) are plotted by study and by population. European ancestry and multi-ancestry random-effects meta-analysis results are plotted as a diamond.


Study





Study
AAGC
CAPPS-ch
SAGE-ch
SAGE-ad
SLSS-ch
SLSJ.ad
SLSJ-ad
FHS
NFBC1966
EPICNCA
EFINIISK-H2000-HBC.YFS FINRISK-H2000-H Chafarge-micic CHAAGE-ARIC
CHARGE-CHS DECODE COPSAC DNBC generation r LISA_GIN maAS
RANE вс58 BCSB
STAMPEED-EUAM CHS-EUAM EGEA-ad egeach ECRHS-ch ECRHS-ch BAMSE BUSSELTON-ch BUSSELTON-ch
BUSSELTON-ad GUBEIIELA-AS KSMU-ch
KSMU-ad MRCA-UK magics-mas PIAMA SAPALDA-ch SAPALDA-ad TOMSK-ch UFA-ch UFA-ch AGARICULTURAL wood BAKERY
ISOCYANATE severe NTR
RS1 RS2 RAG1

European-ancestry
African-ancestry Japanes

Multi-ancestry



11q13.5, rs7927894



Supplementary Figure 4. Quantile-quantile (QQ) plot for the meta-analysis of pediatric asthma QQ plots of $P_{\text {random }}$ values for all SNPs passing QC in at least two-thirds of the studies (27 studies; 8,976 cases, 18,399 controls)

Lambda $=0.951$


## Supplementary Figure 5. Manhattan plot for the meta-analysis of pediatric asthma

The - log10 Prandom values for all SNPs passing QC in at least two-thirds of pediatric studies have been plotted; the dashed horizontal line corresponds to a $P_{\text {random }}$ threshold of $5 \times 10^{-8}$


## Supplementary Figure 6. Regional plots for the nine known asthma loci reaching genome-wide significance

The $x$ axis presents physical distance in megabase (build 37.3) and the $Y$ axis presents $-\log 10 P_{\text {random }}$ values for association statistics in either the multi-ancestry or European ancestry meta-analysis using Locus Zoom. The rs ID is shown for the lead SNP in the region (purple diamond). For remaining SNPs the color indicates $r^{2}$ with the lead SNP.

## 2q12-rs1420101

Multi-ancestry


## European-ancestry



5q22.1 - rs10455025
Multi-ancestry


5q31 - rs20541
Multi-ancestry


## European-ancestry



## European-ancestry



6p21.32 - rs9272346

## Multi-ancestry



9p24.1 - rs992969
Multi-ancestry



European-ancestry


## Multi-ancestry



15 q22.2 - rs11071558
Multi-ancestry



European-ancestry


## Multi-ancestry



17q12-21 - rs2952156

## European-ancestry




## European-ancestry



## Supplementary Figure 7. Regional plots for the 17q12-21 locus in multi-ancestry and pediatric subgroup meta-analyses

The $x$ axis presents physical distance in megabase (build 37.3) and the Y axis presents $-\log 10 P_{\text {random }}$ values for association statistics in either the multi-ancestry or pediatric subgroup meta-analysis using LocusZoom (see URLs). The rs ID is shown for the lead SNP in the region (purple diamond). For remaining SNPs the color indicates $r^{2}$ with the lead SNP.

## Multi-ancestry meta-analysis



Pediatric subgroup meta-analysis


## Supplementary Tables

Supplementary Table 1. Description and study design of the studies included in the meta-analysis

| Consortium Name | Study Name | Study Abbreviation | Ancestry | Country of Residence | Study Design | Number of cases | Number of controls | Pediatric samples ( $P)^{\text {a }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Australian Asthma Genetics Consortium (AAGC) | Australian Asthma Genetics Consortium | AAGC | European | Australia | Population-based and case-control studies | 2,110 | 3,857 |  |
| ALLERGEN | Canadian Asthma Primary Prevention Study (childhood onset) | CAPPS-ch | European | Canada | High-risk birth cohort | 266 | 156 | P |
| ALLERGEN | Canadian Asthma Primary Prevention Study (adult onset) | CAPPS-ad | European | Canada | High-risk birth cohort | 94 | 250 |  |
| ALLERGEN | Study of Asthma Genes and the Environment (childhood onset) | SAGE-ch | European | Canada | Population-based birth cohort | 257 | 267 | P |
| ALLERGEN | Study of Asthma Genes and the Environment (adult onset) | SAGE-ad | European | Canada | Population-based birth cohort | 96 | 402 |  |
| ALLERGEN | Saguenay-Lac-Saint Jean Study (childhood onset) | SLSJ-ch | European | Canada | Family study | 373 | 390 | P |
| ALLERGEN | Saguenay-Lac-Saint Jean Study (adult onset) | SLSJ-ad | European | Canada | Family study | 213 | 223 |  |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | Framingham Heart Study | FHS | European | USA | Population-based cohort | 797 | 6,463 |  |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | Northern Finland Birth Cohort 1966 | NFBC1966 | European | Finland | Population-based birth cohort | 364 | 3,502 |  |
| Analysis in <br> Population-based Cohorts of Asthma Traits (APCAT) | EPIC-Norfolk obese cases | EPIC-N-CA | European | UK | Case Cohort | 123 | 910 |  |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | EPIC-Norfolk population based | EPIC-N-POP | European | UK | Case Cohort | 216 | 2,005 |  |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | FINRISK + Health 2000 (H2000) +Helsinki Birth Cohort (HBC) + Young Finns Study (YFS) | FINRISK- <br> H2000-HBC- <br> YFS | European | Finland | Population-based studies | 555 | 6,923 |  |
| Candidate Gene <br> Association <br> Resource (CARe) <br> Consortium | Atherosclerosis Risk in Communities Study (ARIC) | ARIC-AFAM | African | USA | Population-based study | 148 | 1,087 |  |
| Candidate Gene <br> Association Resource (CARe) Consortium | Corornary Artery Risk Development in Young Adults (CARDIA) | CARDIA | African | USA | Population-based study | 106 | 318 |  |
| Candidate Gene <br> Association Resource (CARe) Consortium | Jackson Heart Study (JHS) | JHS | African | USA | Population-based study | 344 | 1,321 |  |
| Candidate Gene <br> Association <br> Resource (CARe) <br> Consortium | Multi-Ethnic Study of Atherosclerosis (MESA) | MESA-AFAM | African | USA | Population-based study | 294 | 2,275 |  |


| Consortium Name | Study Name | Study Abbreviation | Ancestry | Country of Residence | Study Design | Number of cases | Number of controls | Pediatric samples ( $P)^{\text {a }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Cohort for Heart and Aging Research in Genomic Epidemiology (CHARGE) | Multi-Ethnic Study of Atherosclerosis (MESA) | MESA-EUAM | European | USA | Population-based cohort | 267 | 2,381 | P (for part of the data) |
| Cohort for Heart and Aging Research in Genomic Epidemiology (CHARGE) | Atherosclerosis Risk in Communities Study (ARIC) | ARIC-EUAM | European | USA | Population-based cohort | 453 | 9,713 |  |
| Cohort for Heart and Aging Research in Genomic Epidemiology (CHARGE) | Cardiovascular Health Study (CHS) | CHS | European | USA | Population-based cross sectional study | 179 | 3,058 |  |
| deCODE Genetics | deCODE Genetics | deCODE | European | Iceland | Case-control study | 1,675 | 33,408 |  |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Cophenhagen Study on Asthma in Childdhood | COPSAC | European | Denmark | High-risk birth cohort | 70 | 240 | P |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Danish National Birth Cohort | DNBC | European | Denmark | Population-based birth cohort | 113 | 850 | P |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | GENERATION R | GENERATION R | European | The Netherlands | Population-based birth cohort | 201 | 1,749 | P |
| Early Genetics and <br> Lifecourse <br> Epidemiology <br> (EAGLE) Consortium | LISA Plus \& GINI Plus (German Infant Nutritional Intervention Program) Birth Cohorts | LISA-GINI | European | Germany | Population-based birth cohort | 49 | 999 | P |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Manchester Asthma and Allergy Study | MAAS | European | UK | Population-based birth cohort | 317 | 545 | P |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Western Australian Pregnancy (Raine) Cohort | RAINE | European | Australia | Population-based birth cohort | 328 | 870 | P |
| B58C | British 1958 Birth Cohort | B58C | European | UK | Population-based birth cohort | 986 | 5,505 |  |
| EVE | CAG/CSGA/SARPEUAM | STAMPEEDEUAM | European | USA | Case-control study | 843 | 580 |  |
| EVE | Children'Health Study (CHS) Hispanic White | CHS-LAT | Latino | USA | Case-control study | 606 | 792 | P |
| EVE | CAG/CSGA/SARP-AFAM | STAMPEED- <br> AFAM | African | USA | Case-control study | 644 | 451 |  |
| EVE | Genomic Research on Asthma in the African Diaspora | GRAAD | African | USA | Case-control study | 464 | 471 |  |
| EVE | Study of Asthma <br> Phenotypes and <br> Pharmacogenomics <br> Interactions by Race <br> Ethnicity | SAPPHIRE | African | USA | Case-control study | 149 | 132 |  |
| EVE | Children'Health Study (CHS) Non-Hispanic white | CHS-EUAM | European | USA | Case-control study | 643 | 959 | P |
| Japanese Adult Asthma Research Consortium (JAARC) | Japanese Childhood Onset Asthma in Adult Asthma research | JCOAAD | Japenese | Japan | Case-control study | 301 | 3,304 | P |


| Consortium Name | Study Name | Study Abbreviation | Ancestry | Country of Residence | Study Design | Number of cases | Number of controls | Pediatric samples (P) ${ }^{\text {a }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Japan Pediatric Asthma Consortium (JPAC) | Japan Pediatric Asthma Consortium | JPAC | Japenese | Japan | Case-control study | 938 | 672 | P |
| GABRIEL | Epidemiological study on the Genetics and Environment of Asthma (childhood-onset asthma) | EGEA-ch | European | France | Family study | 482 | 598 | P |
| GABRIEL | Epidemiological study on the Genetics and Environment of Asthma (adult-onset asthma) | EGEA-ad | European | France | Family study | 194 | 654 |  |
| GABRIEL | Avon Longitudinal Study of Parents and Children | ALSPAC | European | United Kingdom | Population-based birth cohort | 607 | 609 | P |
| GABRIEL | European Community Respiratory Health Survey (childhoodonset asthma) | ECRHS-ch | European | Mutliple <br> European <br> Countries | Population-based cohort | 279 | 620 | P |
| GABRIEL | European Community Respiratory Health Survey (adult-onset asthma) | ECRHS-ad | European | Mutliple <br> European Countries | Population-based cohort | 385 | 926 |  |
| GABRIEL | Barn Allergy Milieu Stockholm Epidemiology | BAMSE | European | Sweden | Population-based birth cohort | 239 | 246 | P |
| GABRIEL | Busselton study (childhood-onset asthma) | BUSSELTON-ch | European | Australia | Population-based study | 188 | 390 | P |
| GABRIEL | Busselton study (adultonset asthma) | BUSSELTON-ad | European | Australia | Population-based study | 210 | 419 |  |
| GABRIEL | Gabriel Advanced Surveys | GABRIELA-AS | European | Austria,Germany, Switzerland | Cross-sectional population-based study | 841 | 851 | P |
| GABRIEL | Kursk State Medical University study (childhood-onset asthma) | KSMU-ch | European | Russia | Case-control study | 112 | 116 | P |
| GABRIEL | Kursk State Medical University study (adult-onset asthma) | KSMU-ad | European | Russia | Case-control study | 179 | 161 |  |
| GABRIEL | Medical Research <br> Council Asthma UK <br> Center in Allergic <br> Mechanisms of Asthma <br> (MRCA-UK) | MRCA-UK | European | UK | Family study | 177 | 399 | P |
| GABRIEL | German Multicentre Asthma Genetics in Childhood Study (MAGICS) and Multicentre Allergy Study (MAS) | MAGICS-MAS | European | Germany | Case-control study | 630 | 572 | P |
| GABRIEL | The Prevention and Incidence of Asthma and Mite Allergy birth cohort study | PIAMA | European | The Netherlands | Population-based birth cohort | 172 | 187 | P |
| GABRIEL | Swiss study on Air Pollution And Lung Disease in Adults (childhood-onset asthma) | SAPALDIA-ch | European | Switzerland | Population-based cohort | 237 | 356 | P |
| GABRIEL | Swiss study on Air Pollution And Lung Disease in Adults (adult-onset asthma) | SAPALDIA-ad | European | Switzerland | Population-based cohort | 371 | 557 |  |
| GABRIEL | TOMSK study (childhood-onset asthma) | TOMSK-ch | European | Russia | Family study | 197 | 91 | P |


| Consortium Name | Study Name | Study Abbreviation | Ancestry | Country of Residence | Study Design | Number of cases | Number of controls | Pediatric samples (P) ${ }^{\text {a }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| GABRIEL | TOMSK study (adultonset asthma) | TOMSK-ad | European | Russia | Family study | 62 | 331 |  |
| GABRIEL | UFA region study (childhood-onset asthma) | UFA-ch | European | Russia | Case-control study | 269 | 209 | P |
| GABRIEL | UFA region study (adult-onset asthma) | UFA-ad | European | Russia | Case-control study | 61 | 139 |  |
| GABRIEL | Industrial Cohorts / Agricultural exposures | AGRICULTURAL | European | Mutliple <br> European <br> Countries | Industrial cohort | 256 | 427 |  |
| GABRIEL | Industrial Cohorts / Wood exposure | WOOD | European | Mutliple <br> European Countries | Industrial cohort | 112 | 107 |  |
| GABRIEL | Industrial Cohorts / Wheat and amylase exposures (bakers) | BAKERY | European | Mutliple <br> European Countries | Industrial cohort | 136 | 127 |  |
| GABRIEL | Industrial Cohorts / Isocyanate exposure | ISOCYANATE | European | Mutliple <br> European Countries | Industrial cohort | 25 | 37 |  |
| GABRIEL | Severe asthma study | SEVERE | European | United Kingdom | Case-control study | 290 | 974 |  |
| Netherlands Twin Registry (NTR) | The Netherlands Twin Register | NTR | European | The Netherlands | Twin Registry (Populationbased/Families) | 451 | 2,416 |  |
| Rotterdam Study (RS) | Rotterdam Study 1 | RS1 | European | The Netherlands | Population-based cohort study | 143 | 4,662 |  |
| Rotterdam Study (RS) | Rotterdam Study 2 | RS2 | European | The Netherlands | Population-based cohort study | 67 | 1,616 |  |
| Rotterdam Study (RS) | Rotterdam Study 3 | RS3 | European | The Netherlands | Population-based cohort study | 79 | 1,727 |  |
| Dutch Asthma Genetics Consortium (DAGC) | Dutch Asthma GWAS 1 <br> Family-Based | DAG1 | European | The Netherlands | Family study | 463 | 469 |  |
| Dutch Asthma Genetics Consortium (DAGC) | Dutch Asthma GWAS 2 Case-Control | DAG2 | European | The Netherlands | Case-control study | 452 | 517 |  |
| ${ }^{\text {a Pediatric asthma }}$ is defined as asthma onset $\leq 16$ years of age |  |  |  |  |  |  |  |  |

Supplementary Table 2. Information on genotyping methods, imputation, and statistical analysis by study Details and references for each study can be found in the Supplementary Note

| Consortium Name | Study Name | Study Abbreviation | Genotyping platform | Imputation software | Reference panel used for imputation | Statistical method used for analysis | Software used for analysis |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Australian Asthma Genetics Consortium (AAGC) | Australian Asthma Genetics Consortium | AAGC | Ilumina 370K,610K | IMPUTE2 | $\begin{gathered} \hline \text { HapMap3 (Feb } \\ 2009 \text { release), } \\ 1000 \text { Genomes } \\ \text { (CEU, Mar } 2010 \\ \text { release) } \\ \text { but only } \\ \text { HapMap2 SNPs } \\ \text { provided to TAGC } \end{gathered}$ | Cochran-MantelHaenszel (CMH) test with two strata corresponding to the 610 K and 370 K arrays | PLINK v1.07 |
| ALLERGEN | Canadian Asthma <br> Primary Prevention <br> Study (childhood <br> onset) | CAPPS-ch | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic Regression (robust variance to account for family dependence) | Stata V11 |
| ALLERGEN | Canadian Asthma Primary Prevention Study (adult onset) | CAPPS-ad | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | PLINK |
| ALLERGEN | Study of Asthma Genes and the Environment (childhood onset) | SAGE-ch | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic Regression (robust variance to account for family dependence) | Stata V11 |
| ALLERGEN | Study of Asthma Genes and the Environment (adult onset) | SAGE-ad | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | PLINK |
| ALLERGEN | Saguenay-Lac-Saint Jean Study (childhood onset) | SLSJ-ch | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic Regression (robust variance to account for family dependence) | Stata V10 |
| ALLERGEN | Saguenay-Lac-Saint Jean Study (adult onset) | SLSJ-ad | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic Regression (robust variance to account for family dependence) | Stata V10 |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | Framingham Heart Study | FHS | Affymetrix 5.0 | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | GEE model | GWAF (R <br> package) |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | Northern Finland Birth Cohort 1966 | NFBC1966 | Illumina CNV370 Duo | IMPUTE | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | $\begin{aligned} & \text { QUICKTEST } \\ & \text { v0.94 } \end{aligned}$ |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | EPIC-Norfolk obese cases | EPIC-N-CA | Affymetrix 500K | IMPUTE | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | $\begin{gathered} \text { SNPTEST } \\ 1.1 .5 \end{gathered}$ |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | EPIC-Norfolk population based | EPIC-N-POP | Affymetrix 500K | IMPUTE | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | $\begin{gathered} \text { SNPTEST } \\ 1.1 .5 \end{gathered}$ |
| Analysis in Population-based Cohorts of Asthma Traits (APCAT) | FINRISK + Health 2000 (H2000) <br> +Helsinki Birth Cohort (HBC) + Young Finns Study (YFS) analyzed together | FINRISK | Illumina <br> Human610- Quad; <br> Affymetrix 6.0 | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | $\begin{aligned} & \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |
|  |  | Health 2000 | IlluminaHuman610Quad; Illumina 370K | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | $\begin{aligned} & \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |
|  |  | Helsinki Birth Cohort | Illumina 670K | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \\ \hline \end{gathered}$ | Logistic regression | $\begin{aligned} & \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |
|  |  | Young Finns Syudy | Illumina 670K | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | $\begin{aligned} & \hline \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |


| Consortium Name | Study Name | Study Abbreviation | Genotyping platform | Imputation software | Reference panel used for imputation | Statistical method used for analysis | Software used for analysis |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Candidate Gene <br> Association Resource (CARe) Consortium | Atherosclerosis Risk <br> in Communities <br> Study (ARIC) | ARIC-AFAM | Affymetrix 6.0 | MACH 1.0 | HapMap2 CEU and YRI, r22 | Logistic regression | $\begin{aligned} & \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |
| Candidate Gene <br> Association Resource (CARe) Consortium | Corornary Artery Risk Development in Young Adults (CARDIA) | CARDIA | Affymetrix 6.0 | MACH 1.0 | $\begin{aligned} & \text { HapMap2 CEU } \\ & \text { and YRI, r22 } \end{aligned}$ | Logistic regression | $\begin{aligned} & \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |
| Candidate Gene <br> Association Resource (CARe) Consortium | Jackson Heart Study (JHS) | JHS | Affymetrix 6.0 | MACH 1.0 | $\begin{aligned} & \text { HapMap2 CEU } \\ & \text { and YRI, r22 } \end{aligned}$ | Logistic regression | $\begin{aligned} & \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |
| Candidate Gene <br> Association <br> Resource (CARe) <br> Consortium | Multi-Ethnic Study of Atherosclerosis (MESA) | MESA-AFAM | Affymetrix 6.0 | IMPUTE2 | HapMap2 (all populations), r22 | Additive model using GEE to account for familial correlation | GWAF (R <br> package) |
| Cohort for Heart and Aging Research in Genomic Epidemiology (CHARGE) | Multi-Ethnic Study of Atherosclerosis (MESA) | MESA-EUAM | Affymetrix 6.0 | IMPUTE2 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r24 } \end{gathered}$ | Logistic regression (frequentist association test) | $\begin{gathered} \text { SNPTEST } \\ \text { v2.2.0 } \end{gathered}$ |
| Cohort for Heart and Aging Research in Genomic Epidemiology (CHARGE) | Atherosclerosis Risk in Communities Study (ARIC) | ARIC-EUAM | Affymetrix 6.0 | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | $\begin{gathered} \text { ProbABEL } \\ \text { v.0.1-3 } \end{gathered}$ |
| Cohort for Heart and Aging Research in Genomic Epidemiology (CHARGE) | Cardiovascular Health Study (CHS) | CHS | Illumina 370K CNV | BIMBAM | HapMap CEU, r22 | Logistic regression with robust SE estimates | R |
| decCODE Genetics | deCODE Genetics | deCODE | Illumina HumanHap300 or HumanHapCNV370 | IMPUTE | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | SNPTEST v2 |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Cophenhagen Study on Asthma in Childdhood | COPSAC | Illumina 550K | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | $\begin{gathered} \text { SNPTEST } \\ \text { v2.2.0 } \end{gathered}$ |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Danish National Birth Cohort | DNBC | Illumina 660K | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | PLINK v1.07 <br> MACH2DAT |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | GENERATION R | GENERATION R | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | GRIMP |
| Early Genetics and <br> Lifecourse <br> Epidemiology <br> (EAGLE) Consortium | LISA Plus \& GINI Plus (German Infant Nutritional Intervention Program) Birth Cohorts | LISA-GINI | Affymetrix 5.0 and $6.0$ | IMPUTE2 | Hapmap2, r22 | Logistic regression | $\begin{gathered} \text { SNPTEST } \\ \text { v2.2.0 } \end{gathered}$ |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Manchester Asthma and Allergy Study | MAAS | Illumina <br> Human610-Quad | IMPUTE2 | HapMap 3 + 1,000 Genomes | Logistic regression | $\begin{gathered} \text { SNPTEST } \\ \text { v2.4.0 } \end{gathered}$ |
| Early Genetics and Lifecourse Epidemiology (EAGLE) Consortium | Western Australian Pregnancy (Raine) Cohort | RAINE | Illumina 660K | MACH 1.0., Minimac | HapMap2 r22 | Logistic regression | $\begin{gathered} \text { SNPTEST } \\ \text { v2.2.0 } \end{gathered}$ |


| Consortium Name | Study Name | Study Abbreviation | Genotyping platform | Imputation software | Reference panel used for imputation | Statistical method used for analysis | Software used for analysis |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| B58C | British 1958 Birth Cohort | B58C | Illumina 550K and Illumina Human 610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | ProbABEL $0.1-3$ <br> (palogist) |
| EVE | CAG/CSGA/SARP- EUAM | STAMPEEDEUAM | Illumina 1Mv1 | MACH 1.0 | $\begin{gathered} \hline \text { HapMap2 CEU, } \\ \text { r21 } \\ \hline \end{gathered}$ | Logistic regression | R |
| EVE | Children'Health Study (CHS) Hispanic White | CHS-LAT | Illumina 550K and Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 } \\ \text { CEU+ASN, r21 } \end{gathered}$ | Logistic regression | R |
| EVE | $\begin{aligned} & \text { CAG/CSGA/SARP- } \\ & \text { AFAM } \end{aligned}$ | STAMPEEDAFAM | Illumina 1Mv1 | MACH 1.0 | $\begin{gathered} \text { HapMap2 } \\ \text { CEU+YRI, r21 } \end{gathered}$ | Logistic regression | R |
| EVE | Genomic Research on Asthma in the African Diaspora | GRAAD | Illumina 650K | MACH 1.0 | $\begin{gathered} \text { HapMap2 } \\ \text { CEU+YRI+ASN, } \\ \text { r21 } \\ \hline \end{gathered}$ | T-test | R |
| EVE | Study of Asthma Phenotypes and Pharmacogeneomics Interactions by Race Ethnicity | SAPPHIRE | Affymetrix 5.0, 6.0 and mapping 500k array | MACH 1.0 | $\begin{gathered} \text { HapMap2 } \\ \text { CEU+YRI, r21 } \end{gathered}$ | Logistic regression | R |
| EVE | Children'Health Study (CHS) NonHispanic white | CHS-EUAM | Illumina 550 K and Illumina Human 610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | R |
| Japanese Adult Asthma Research Consortium (JAARC) | Japanese Childhood Onset Asthma in Adult Asthma research | JCOAAD | Illumina 550K and Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 } \\ \text { JPT+CHB, r21 } \end{gathered}$ | Logistic regression | MACH2DAT |
| Japan Pediatric Asthma Consortium (JPAC) | Japan Pediatric <br> Asthma Consortium | JPAC | Illumina 550K and Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 } \\ \text { JPT+CHB, r21 } \end{gathered}$ | Logistic regression | MACH2DAT |
| GABRIEL | Epidemiological study on the Genetics and Environment of Asthma (childhoodonset asthma) | EGEA-ch | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression (robust variance to account for family dependence) | Stata V10 |
| GABRIEL | Epidemiological <br> study on the <br> Genetics and <br> Environment of <br> Asthma (adult-onset <br> asthma) | EGEA-ad | Illumina Human610-Quad | MACH 1.0 | HapMap2 CEU, r21 | Logistic regression (robust variance to account for family dependence) | Stata V10 |
| GABRIEL | Avon Longitudinal Study of Parents and Children | ALSPAC | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | European <br> Community <br> Respiratory Health <br> Survey (childhood- <br> onset asthma) | ECRHS-ch | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | European <br> Community <br> Respiratory Health <br> Survey (adult-onset <br> asthma) | ECRHS-ad | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Barn Allergy Milieu <br> Stockholm <br> Epidemiology | BAMSE | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Busselton study (childhood-onset asthma) | BUSSELTON-ch | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Busselton study (adult-onset asthma) | BUSSELTON-ad | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |


| Consortium Name | Study Name | Study Abbreviation | Genotyping platform | Imputation software | Reference panel used for imputation | Statistical method used for analysis | Software used for analysis |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| GABRIEL | Gabriel Advanced Surveys | GABRIELA-AS | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \\ \hline \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Kursk State Medical University study (childhood-onset asthma) | KSMU-ch | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Kursk State Medical University study (adult-onset asthma) | KSMU-ad | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Medical Research <br> Council Asthma UK <br> Center in Allergic <br> Mechanisms of <br> Asthma (MRCA-UK) | MRCA-UK | Illumina Sentrix <br> Human-1 and <br> Sentrix <br> HumanHap300 | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression (robust variance to account for family dependence) | Stata V10 |
| GABRIEL | German Multicentre Asthma Genetics in Childhood Study (MAGICS) and Multicentre Allergy Study (MAS) | MAGICS-MAS | Illumina Sentrix <br> HumanHap300 <br> (MAGICS) <br> Illumina <br> Human610-Quad (MAS) | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | The Prevention and Incidence of Asthma and Mite Allergy birth cohort study | PIAMA | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Swiss study on Air Pollution And Lung Disease in Adults (childhood-onset asthma) | SAPALDIA-ch | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Swiss study on Air Pollution And Lung Disease in Adults (adult-onset asthma) | SAPALDIA-ad | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | TOMSK study (childhood-onset asthma) | TOMSK-ch | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression (robust variance to account for family dependence) | Stata V10 |
| GABRIEL | TOMSK study (adultonset asthma) | TOMSK-ad | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression (robust variance to account for family dependence) | Stata V10 |
| GABRIEL | UFA region study (childhood-onset asthma) | UFA-ch | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | UFA region study (adult-onset asthma) | UFA-ad | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Industrial Cohorts / Agricultural exposures | AGRICULTURAL | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Industrial Cohorts / Wood exposure | WOOD | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Industrial Cohorts / Wheat and amylase exposures (bakers) | BAKERY | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Industrial Cohorts / Isocyanate exposure | ISOCYANATE | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| GABRIEL | Severe asthma study | SEVERE | Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r21 } \end{gathered}$ | Logistic regression | Stata V10 |
| Netherlands Twin Registry (NTR) | The Netherlands Twin Register | NTR | Illumina 660K, Illumina Omni Express 1 mil, Affymetrix Perlegen and Affymetrix 6.0 | IMPUTE2 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r24 } \end{gathered}$ | Logistic regression | $\begin{aligned} & \text { PLINK } \\ & \text { v1.07 } \end{aligned}$ |


| Consortium Name | Study Name | Study Abbreviation | Genotyping platform | Imputation software | Reference panel used for imputation | Statistical method used for analysis | Software used for analysis |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Rotterdam Study (RS) | Rotterdam Study 1 | RS1 | Illumina 550K | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \\ \hline \end{gathered}$ | Logistic regression | MACH2DAT (GRIMP) |
| Rotterdam Study (RS) | Rotterdam Study 2 | RS2 | Illumina 550K and Illumina Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | $\begin{aligned} & \text { MACH2DAT } \\ & \text { (GRIMP) } \end{aligned}$ |
| Rotterdam Study (RS) | Rotterdam Study 3 | RS3 | Illumina <br> Human610-Quad | MACH 1.0 | $\begin{gathered} \text { HapMap2 CEU, } \\ \text { r22 } \\ \hline \end{gathered}$ | Logistic regression | $\begin{aligned} & \text { MACH2DAT } \\ & \text { (GRIMP) } \\ & \hline \end{aligned}$ |
| Dutch Asthma Genetics Consortium (DAGC) | Dutch Asthma GWAS 1 FamilyBased | DAG1 | Illumina 317K | BEAGLE | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | PLINK v1.07 |
| Dutch Asthma Genetics Consortium (DAGC) | Dutch Asthma GWAS 2 CaseControl | DAG2 | Illumina 370 Duo | BEAGLE | $\begin{gathered} \text { Hapmap2 CEU, } \\ \text { r22 } \end{gathered}$ | Logistic regression | PLINK v1.07 |

Supplementary Table 3. Effect sizes of the lead SNPs in ancestry-specific and multi-ancestry meta-analyses
The lead SNPs belong to the 18 loci that are genome-wide significant ( $P_{\text {random }} \leq 10^{-8}$ ) in European-ancestry and/or multi-ancestry meta-analyse (as shown in Table 1)

| Region ${ }^{\text {a }}$ | SNP | Position | Allele (R/E) ${ }^{\text {c }}$ | EUROPEAN ANCESTRY (56 studies) |  |  |  |  | AFRICAN ANCESTRY (7 studies) |  |  |  | JAPANESE (2 studies) |  |  |  | LATINOS (1 study) |  |  | MULTTI-ANCESTRY (66 studies) |  |  |  | $\mathrm{P}_{\text {etmic }}{ }^{\text {b }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  |  | EAF ${ }^{\text {d }}$ | OR ( $95 \% \mathrm{Cl})^{\text {e }}$ | Prandom ${ }^{\text {f }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {b }}$ | $\mathrm{P}_{\text {net }}{ }^{\text {n }}$ | EAF | OR ( $95 \% \mathrm{Cl})^{\text {e }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {b }}$ | $\mathrm{P}_{\text {net }}{ }^{\text {n }}$ | EAF | OR (95\% CI) ${ }^{\text {e }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {g }}$ | $\mathrm{P}_{\text {net }}{ }^{\text {n }}$ | EAF | OR ( $95 \% \mathrm{Cl})^{\text {e }}$ | $\mathrm{P}^{\text {i }}$ | OR (95\% CI) ${ }^{\text {e }}$ | $\mathrm{Prandom}^{\text {f }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {g }}$ | $\mathrm{P}_{\text {net }}{ }^{\text {n }}$ |  |


| New |  | oci |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 5 q 31.3 | rs7705042 | 141,492,419 | C/A | 0.63 | 1.08 (1.05-1.11) | $1.6 \times 10^{-6}$ | $8.5 \times 10^{-10}$ | 0.07 | 0.77 | 1.12 (1.0-1.24) | 0.04 | 0.39 | 0.63 | 1.17 (1.05-1.31) | $5.5 \times 10^{-3}$ | 0.98 | 0.67 | 1.10 (0.93-1.3) | 0.27 | 1.09 (1.06-1.12) | $7.9 \times 10^{-9}$ | $3.1 \times 10^{-12} 0.11$ | 0.53 |
| 6p22.1 | rs1233578 | 28,712,247 | A/G | 0.13 | 1.11 (1.07-1.15) | $5.3 \times 10^{-9}$ | $5.3 \times 10^{-9}$ | 0.82 | 0.41 | 0.98 (0.90-1.07) | 0.65 | 0.73 |  | NA | NA | NA | 0.15 | 0.83 (0.66-1.05) | 0.12 | 1.09 (1.05-1.12) | $5.9 \times 10^{-7}$ | $5.9 \times 10^{-7} 0.56$ | 0.003 |
| 6 q 15 | rs2325291 | 90,986,686 | G/A | 0.33 | 0.91 (0.89-0.93) | $8.6 \times 10^{-13}$ | $8.6 \times 10^{-13}$ | 0.78 | 0.12 | 1.02 (0.89-1.17) | 0.78 | 0.49 | 0.03 | 1.02 (0.74-1.41) | 0.88 | 0.69 | 0.30 | 0.91 (0.76-1.09) | 0.30 | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | $2.2 \times 10^{-12} 0.8$ | 0.39 |
| 12q13.3 | rs167769 | 57,503,775 | C/T | 0.4 | 1.08 (1.05-1.11) | $1.6 \times 10^{-7}$ | $5.5 \times 10^{-9}$ | 0.19 | 0.14 | 1.14 (1.0-1.29) | 0.05 | 0.71 | 0.21 | 1.09 (0.95-1.24) | 0.22 | 0.18 | 0.37 | 1.04 (0.89-1.23) | 0.60 | 1.08 (1.05-1.11) | $3.9 \times 10^{-9}$ | $4.6 \times 10^{-10} 0.31$ | 0.87 |
| 17q21.33 | rs17637472 | 47,461,433 | G/A | 0.39 | 1.08 (1.05-1.11) | $3.3 \times 10^{-9}$ | $3.3 \times 10^{-9}$ | 0.56 | 0.08 | 0.91 (0.77-1.09) | 0.32 | 0.13 | 0.13 | 1.12 (0.95-1.32) | 0.18 | 0.58 | 0.22 | 1.24 (1.03-1.5) | 0.02 | 1.08 (1.05-1.11) | $6.6 \times 10^{-9}$ | $1.1 \times 10^{-9} 0.35$ | 0.12 |

## New signals at loci previously assosiated with asthma in ancestry-specific populations



## Asthma signals previously reported for asthma plus hay fever




## Knwon asthma loci

$2 q 12$ rs1420101 102,957,716
5q22.1 rs10455025 110,404,999
5q31 rs20541 131,995,964
A/G $\begin{array}{llll}0.79 & 0.89 & (0.86-0.91) & 1.4 \times 10^{-14} \\ 1.4 \times 10^{-14} & 0.73\end{array}$
G/A $\left\lvert\, \begin{array}{llll}0.56 & 1.16(1.13-1.19) & 4.8 \times 10^{-28} & 2.4 \times 10^{-28} \\ 0.46\end{array}\right.$
11q13.5 rs7927894 76,301,316
15q22.2 rs11071558 61,069,421
15q22.33 rs2033784 67,449,660 A/G $\left\lvert\, \begin{array}{llllll} & 1.11(1.08-1.14) & 2.5 \times 10^{-14} 2.5 \times 10^{-14} & 0.75\end{array}\right.$

${ }^{2}$ Cytogenetic position of the chromosomal region harboring the lead SNP
${ }^{\mathrm{b}}$ Genes coding for proteins; the gene where eventually the lead SNP lies is first indicated followed by the previous gene and next gene
${ }^{\mathrm{C}} \mathrm{R}=$ reference allele $/ \mathrm{E}=$ effect allele. ${ }^{\mathrm{d}} \mathrm{EAF}=$ Effect allele frequency

${ }^{\mathrm{f}} \mathrm{Prandom}$ is P -value for test of association between SNP and asthma under a random-effects model
${ }^{\mathrm{E}} \mathrm{P}_{\text {fixed }}$ is P -value for test of association between SNP and asthma under a fixed-effects model
${ }^{h} P_{\text {het }}$ is the $P$-value for test of heterogeneity across studies with the use of Cochran's $Q$ test
${ }^{\text {i }} \mathrm{P}$ for test of association between SNP and asthma in the Latino GWAS
${ }^{\mathrm{j}} \mathrm{P}_{\text {ethnic }}$ for test of heterogeneity between ancestry-specific effect sizes
$N A=$ the SNP did not pass QC in the Japanese studies

Supplementary Table 4. SNPs with $P_{\text {random }} \leq 10^{-8}$ in the European ancestry meta-analysis

| Chrom | SNP | Position | Allele <br> (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{\mathrm{b}} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{gathered} \text { OR }_{\text {fixed }} \\ (95 \% \mathrm{Cl})^{b} \end{gathered}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 | rs1465325 | 102,734,551 | A/G | 0.17 | 0.88 (0.85-0.92) | $5.09 \times 10^{-10}$ | 0.88 (0.85-0.91) | $3.68 \times 10^{-14}$ | 0.12 | 100\% |
| 2 | rs1465324 | 102,735,612 | G/A | 0.18 | 0.89 (0.86-0.93) | $2.71 \times 10^{-9}$ | 0.89 (0.86-0.92) | $7.69 \times 10^{-13}$ | 0.16 | 100\% |
| 2 | rs12328681 | 102,736,867 | G/T | 0.18 | 0.89 (0.86-0.93) | $2.83 \times 10^{-9}$ | 0.89 (0.86-0.92) | $7.19 \times 10^{-13}$ | 0.15 | 100\% |
| 2 | rs10172039 | 102,738,785 | C/A | 0.18 | 0.89 (0.86-0.93) | $2.72 \times 10^{-9}$ | 0.89 (0.86-0.92) | $7.39 \times 10^{-13}$ | 0.15 | 100\% |
| 2 | rs12619383 | 102,741,718 | A/G | 0.17 | 0.88 (0.85-0.92) | $1.67 \times 10^{-9}$ | 0.88 (0.85-0.91) | $5.95 \times 10^{-14}$ | 0.09 | 100\% |
| 2 | rs12620132 | 102,742,115 | A/G | 0.17 | 0.88 (0.85-0.92) | $1.12 \times 10^{-9}$ | 0.88 (0.85-0.91) | $3.88 \times 10^{-14}$ | 0.09 | 100\% |
| 2 | rs13392285 | 102,744,338 | C/A | 0.18 | 0.89 (0.86-0.92) | $4.72 \times 10^{-10}$ | 0.89 (0.86-0.91) | $2.12 \times 10^{-13}$ | 0.19 | 100\% |
| 2 | rs13387400 | 102,750,020 | G/A | 0.17 | 0.88 (0.85-0.92) | $1.16 \times 10^{-9}$ | 0.88 (0.85-0.91) | $4.37 \times 10^{-14}$ | 0.09 | 100\% |
| 2 | rs1558641 | 102,765,865 | G/A | 0.17 | 0.88 (0.85-0.92) | $7.85 \times 10^{-9}$ | 0.88 (0.85-0.91) | $6.20 \times 10^{-14}$ | 0.05 | 100\% |
| 2 | rs1558640 | 102,765,878 | G/A | 0.17 | 0.88 (0.85-0.92) | $4.12 \times 10^{-9}$ | 0.88 (0.85-0.91) | $4.73 \times 10^{-14}$ | 0.06 | 100\% |
| 2 | rs871657 | 102,771,341 | C/T | 0.18 | 0.89 (0.85-0.93) | $9.31 \times 10^{-9}$ | 0.88 (0.86-0.91) | $9.67 \times 10^{-14}$ | 0.07 | 100\% |
| 2 | rs2287048 | 102,773,999 | C/T | 0.17 | 0.88 (0.85-0.92) | $2.45 \times 10^{-9}$ | 0.88 (0.85-0.91) | $3.50 \times 10^{-14}$ | 0.06 | 100\% |
| 2 | rs1997502 | 102,844,249 | A/G | 0.65 | 1.09 (1.06-1.13) | $6.51 \times 10^{-9}$ | 1.09 (1.06-1.12) | $3.20 \times 10^{-10}$ | 0.26 | 96\% |
| 2 | rs10189629 | 102,879,464 | C/A | 0.14 | 0.85 (0.81-0.89) | $2.16 \times 10^{-10}$ | 0.85 (0.82-0.89) | $1.49 \times 10^{-15}$ | 0.02 | 98\% |
| 2 | rs11692065 | 102,883,975 | C/T | 0.14 | 0.85 (0.81-0.89) | $1.47 \times 10^{-10}$ | 0.85 (0.82-0.89) | $1.21 \times 10^{-15}$ | 0.02 | 98\% |
| 2 | rs11674302 | 102,887,128 | T/C | 0.14 | 0.85 (0.81-0.89) | $8.05 \times 10^{-11}$ | 0.85 (0.82-0.89) | $7.92 \times 10^{-16}$ | 0.03 | 98\% |
| 2 | rs11690644 | 102,914,214 | A/G | 0.14 | 0.84 (0.8-0.88) | $7.74 \times 10^{-13}$ | 0.84 (0.81-0.87) | $6.93 \times 10^{-20}$ | 0.03 | 96\% |
| 2 | rs950880 | 102,932,562 | C/A | 0.38 | 1.12 (1.09-1.15) | $3.95 \times 10^{-19}$ | 1.12 (1.09-1.15) | $3.95 \times 10^{-19}$ | 0.55 | 100\% |
| 2 | rs13001325 | 102,939,036 | C/T | 0.38 | 1.12 (1.09-1.15) | $2.02 \times 10^{-19}$ | 1.12 (1.09-1.15) | $2.02 \times 10^{-19}$ | 0.55 | 100\% |
| 2 | rs12479210 | 102,949,161 | C/T | 0.38 | 1.12 (1.09-1.15) | $2.14 \times 10^{-19}$ | 1.12 (1.09-1.15) | $2.14 \times 10^{-19}$ | 0.57 | 100\% |
| 2 | rs13019081 | 102,950,822 | A/C | 0.38 | 1.12 (1.09-1.15) | $1.99 \times 10^{-19}$ | 1.12 (1.09-1.15) | $1.99 \times 10^{-19}$ | 0.57 | 100\% |
| 2 | rs17026974 | 102,952,360 | G/A | 0.29 | 1.1 (1.08-1.13) | $2.30 \times 10^{-13}$ | 1.1 (1.08-1.13) | $2.30 \times 10^{-13}$ | 0.62 | 100\% |
| 2 | rs3771180 | 102,953,617 | G/T | 0.14 | 0.84 (0.8-0.88) | $1.11 \times 10^{-13}$ | 0.84 (0.81-0.87) | $1.47 \times 10^{-20}$ | 0.04 | 100\% |
| 2 | rs13431828 | 102,954,653 | C/T | 0.14 | 0.84 (0.8-0.88) | $9.10 \times 10^{-14}$ | 0.84 (0.81-0.87) | $1.14 \times 10^{-19}$ | 0.07 | 100\% |
| 2 | rs13408661 | 102,955,082 | G/A | 0.14 | 0.84 (0.8-0.88) | $1.33 \times 10^{-13}$ | 0.84 (0.81-0.87) | $5.52 \times 10^{-20}$ | 0.05 | 100\% |
| 2 | rs873022 | 102,955,683 | G/T | 0.29 | 1.11 (1.08-1.13) | $1.57 \times 10^{-13}$ | 1.11 (1.08-1.13) | $1.57 \times 10^{-13}$ | 0.63 | 100\% |
| 2 | rs3771177 | 102,955,860 | G/T | 0.29 | 1.11 (1.08-1.13) | $1.53 \times 10^{-13}$ | 1.11 (1.08-1.13) | $1.53 \times 10^{-13}$ | 0.64 | 100\% |
| 2 | rs10173081 | 102,957,348 | C/T | 0.14 | 0.84 (0.8-0.88) | $1.43 \times 10^{-13}$ | 0.84 (0.81-0.87) | $6.85 \times 10^{-20}$ | 0.05 | 100\% |
| 2 | rs3732129 | 102,957,532 | T/C | 0.29 | 1.1 (1.08-1.13) | $1.61 \times 10^{-13}$ | 1.1 (1.08-1.13) | $1.61 \times 10^{-13}$ | 0.65 | 100\% |
| 2 | rs1420101 | 102,957,716 | C/T | 0.37 | 1.12 (1.1-1.15) | $9.08 \times 10^{-20}$ | 1.12 (1.1-1.15) | $9.08 \times 10^{-20}$ | 0.63 | 100\% |
| 2 | rs12905 | 102,960,007 | G/A | 0.29 | 1.1 (1.08-1.13) | $2.21 \times 10^{-13}$ | 1.1 (1.08-1.13) | $2.21 \times 10^{-13}$ | 0.66 | 100\% |
| 2 | rs12712142 | 102,960,584 | C/A | 0.40 | 1.12 (1.09-1.15) | $2.87 \times 10^{-17}$ | 1.12 (1.09-1.15) | $3.93 \times 10^{-19}$ | 0.39 | 98\% |
| 2 | rs2160203 | 102,960,824 | A/G | 0.24 | 0.89 (0.86-0.92) | $5.89 \times 10^{-13}$ | 0.89 (0.86-0.92) | $5.89 \times 10^{-13}$ | 0.67 | 89\% |
| 2 | rs13017455 | 102,964,742 | C/T | 0.39 | 1.12 (1.09-1.15) | $2.66 \times 10^{-18}$ | 1.12 (1.09-1.15) | $3.12 \times 10^{-19}$ | 0.43 | 100\% |
| 2 | rs12469506 | 102,965,871 | C/T | 0.30 | 1.12 (1.09-1.15) | $2.86 \times 10^{-15}$ | 1.12 (1.09-1.15) | $2.86 \times 10^{-15}$ | 0.52 | 96\% |
| 2 | rs1921622 | 102,966,067 | G/A | 0.53 | 1.09 (1.07-1.12) | $6.14 \times 10^{-13}$ | 1.09 (1.07-1.12) | $6.14 \times 10^{-13}$ | 0.50 | 100\% |
| 2 | rs10208293 | 102,966,310 | G/A | 0.28 | 0.88 (0.85-0.92) | $8.00 \times 10^{-10}$ | 0.88 (0.86-0.91) | $9.42 \times 10^{-19}$ | 0.00 | 100\% |
| 2 | rs10197862 | 102,966,549 | A/G | 0.14 | 0.84 (0.8-0.88) | $1.17 \times 10^{-12}$ | 0.84 (0.81-0.88) | $2.23 \times 10^{-19}$ | 0.03 | 100\% |
| 2 | rs1861245 | 102,966,906 | C/T | 0.38 | 0.9 (0.87-0.93) | $5.33 \times 10^{-12}$ | 0.9 (0.88-0.93) | $4.06 \times 10^{-15}$ | 0.18 | 100\% |
| 2 | rs13424006 | 102,967,236 | T/C | 0.39 | 0.89 (0.87-0.92) | $9.42 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.59 \times 10^{-18}$ | 0.15 | 100\% |
| 2 | rs6751967 | 102,967,413 | T/C | 0.39 | 0.89 (0.87-0.92) | $9.01 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.40 \times 10^{-18}$ | 0.15 | 100\% |
| 2 | rs6749114 | 102,967,587 | A/C | 0.39 | 0.89 (0.87-0.92) | $8.61 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.98 \times 10^{-18}$ | 0.16 | 100\% |
| 2 | rs11123923 | 102,967,844 | C/A | 0.39 | 1.12 (1.09-1.15) | $6.03 \times 10^{-18}$ | 1.12 (1.09-1.15) | $1.59 \times 10^{-19}$ | 0.40 | 100\% |
| 2 | rs4988955 | 102,967,928 | A/G | 0.39 | 0.89 (0.87-0.92) | $7.24 \times 10^{-14}$ | 0.89 (0.87-0.92) | $3.32 \times 10^{-18}$ | 0.17 | 100\% |
| 2 | rs4988956 | 102,968,007 | G/A | 0.39 | 0.89 (0.87-0.92) | $4.41 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.00 \times 10^{-18}$ | 0.17 | 100\% |
| 2 | rs4988957 | 102,968,075 | T/C | 0.39 | 0.89 (0.87-0.92) | $7.22 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.55 \times 10^{-18}$ | 0.16 | 100\% |
| 2 | rs10204137 | 102,968,212 | A/G | 0.39 | 0.89 (0.86-0.92) | $3.08 \times 10^{-12}$ | 0.89 (0.87-0.92) | $1.49 \times 10^{-16}$ | 0.13 | 91\% |
| 2 | rs4988958 | 102,968,285 | T/C | 0.39 | 0.89 (0.87-0.92) | $7.07 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.43 \times 10^{-18}$ | 0.16 | 100\% |
| 2 | rs10192157 | 102,968,356 | C/T | 0.39 | 0.89 (0.87-0.92) | $6.33 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.90 \times 10^{-18}$ | 0.17 | 100\% |
| 2 | rs10206753 | 102,968,362 | T/C | 0.39 | 0.89 (0.86-0.92) | $2.16 \times 10^{-12}$ | 0.89 (0.87-0.92) | $1.29 \times 10^{-16}$ | 0.14 | 91\% |
| 2 | rs7603730 | 102,974,371 | A/C | 0.39 | 0.89 (0.87-0.92) | $6.72 \times 10^{-14}$ | 0.89 (0.87-0.92) | $1.97 \times 10^{-18}$ | 0.16 | 100\% |
| 2 | rs12998521 | 102,974,417 | G/T | 0.39 | 1.12 (1.09-1.15) | $1.98 \times 10^{-15}$ | 1.12 (1.09-1.15) | $6.77 \times 10^{-18}$ | 0.33 | 98\% |
| 2 | rs10170583 | 102,974,764 | G/A | 0.39 | 0.89 (0.87-0.92) | $2.81 \times 10^{-13}$ | 0.89 (0.87-0.92) | $2.71 \times 10^{-18}$ | 0.12 | 100\% |
| 2 | rs10176664 | 102,976,172 | G/A | 0.38 | 0.89 (0.87-0.92) | $7.43 \times 10^{-14}$ | 0.89 (0.87-0.92) | $1.90 \times 10^{-18}$ | 0.16 | 100\% |
| 2 | rs3755276 | 102,978,459 | C/T | 0.38 | 0.89 (0.87-0.92) | $8.01 \times 10^{-14}$ | 0.89 (0.87-0.92) | $1.96 \times 10^{-18}$ | 0.15 | 100\% |
| 2 | rs2287037 | 102,979,028 | C/T | 0.39 | 1.12 (1.09-1.15) | $2.41 \times 10^{-17}$ | 1.12 (1.09-1.15) | $1.19 \times 10^{-19}$ | 0.36 | 100\% |


| Chrom | SNP | Position | Allele <br> (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $P_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 | rs3771172 | 102,985,812 | C/T | 0.29 | 1.11 (1.08-1.14) | $1.24 \times 10^{-14}$ | 1.11 (1.08-1.14) | $1.24 \times 10^{-14}$ | 0.48 | 100\% |
| 2 | rs3771171 | 102,985,950 | T/C | 0.29 | 1.11 (1.08-1.14) | $1.05 \times 10^{-13}$ | 1.11 (1.08-1.14) | $2.01 \times 10^{-14}$ | 0.42 | 96\% |
| 2 | rs2160202 | 102,986,154 | G/A | 0.29 | 1.11 (1.08-1.14) | $1.12 \times 10^{-13}$ | 1.11 (1.08-1.14) | $2.00 \times 10^{-14}$ | 0.42 | 96\% |
| 2 | rs3771166 | 102,986,222 | G/A | 0.38 | 0.89 (0.86-0.91) | $7.64 \times 10^{-18}$ | 0.89 (0.86-0.91) | $1.31 \times 10^{-19}$ | 0.38 | 95\% |
| 2 | rs1974675 | 102,986,375 | G/A | 0.38 | 0.89 (0.86-0.91) | $7.47 \times 10^{-18}$ | 0.89 (0.86-0.91) | $1.26 \times 10^{-19}$ | 0.38 | 95\% |
| 2 | rs10439410 | 102,990,788 | G/T | 0.49 | 0.92 (0.89-0.94) | $1.49 \times 10^{-9}$ | 0.92 (0.9-0.94) | $2.21 \times 10^{-11}$ | 0.22 | 100\% |
| 2 | rs6758936 | 102,991,369 | G/A | 0.49 | 0.92 (0.89-0.94) | $1.24 \times 10^{-9}$ | 0.92 (0.9-0.94) | $2.29 \times 10^{-11}$ | 0.23 | 100\% |
| 2 | rs2041739 | 102,994,333 | C/T | 0.49 | 0.92 (0.89-0.94) | $9.48 \times 10^{-10}$ | 0.92 (0.9-0.94) | $2.32 \times 10^{-11}$ | 0.25 | 100\% |
| 2 | rs17027037 | 102,994,884 | A/G | 0.29 | 1.11 (1.08-1.14) | $4.15 \times 10^{-14}$ | 1.11 (1.08-1.14) | $4.15 \times 10^{-14}$ | 0.51 | 100\% |
| 2 | rs2080289 | 102,995,020 | G/A | 0.29 | 1.11 (1.08-1.14) | $4.61 \times 10^{-14}$ | 1.11 (1.08-1.14) | $4.61 \times 10^{-14}$ | 0.52 | 100\% |
| 2 | rs10208196 | 102,996,345 | G/A | 0.49 | 0.92 (0.89-0.94) | $9.18 \times 10^{-10}$ | 0.92 (0.9-0.94) | $2.56 \times 10^{-11}$ | 0.26 | 100\% |
| 2 | rs11683700 | 102,996,805 | C/T | 0.29 | 1.11 (1.08-1.14) | $7.04 \times 10^{-14}$ | 1.11 (1.08-1.14) | $6.65 \times 10^{-14}$ | 0.47 | 96\% |
| 2 | rs3213732 | 102,998,279 | A/G | 0.49 | 0.92 (0.89-0.94) | $8.97 \times 10^{-10}$ | 0.92 (0.9-0.94) | $2.38 \times 10^{-11}$ | 0.25 | 100\% |
| 2 | rs6760621 | 102,999,952 | T/C | 0.49 | 0.92 (0.89-0.94) | $2.94 \times 10^{-10}$ | 0.92 (0.89-0.94) | $3.53 \times 10^{-11}$ | 0.36 | 91\% |
| 2 | rs1035130 | 103,001,402 | C/T | 0.29 | 1.11 (1.08-1.14) | $4.15 \times 10^{-14}$ | 1.11 (1.08-1.14) | $4.15 \times 10^{-14}$ | 0.53 | 100\% |
| 2 | rs6706002 | 103,006,104 | A/G | 0.49 | 0.92 (0.89-0.94) | $8.57 \times 10^{-10}$ | 0.92 (0.9-0.94) | $2.17 \times 10^{-11}$ | 0.25 | 100\% |
| 2 | rs4851570 | 103,006,387 | A/G | 0.29 | 1.11 (1.08-1.14) | $3.27 \times 10^{-14}$ | 1.11 (1.08-1.14) | $3.27 \times 10^{-14}$ | 0.52 | 100\% |
| 2 | rs6749014 | 103,006,448 | C/T | 0.49 | 0.92 (0.89-0.94) | $1.15 \times 10^{-9}$ | 0.92 (0.9-0.94) | $4.83 \times 10^{-11}$ | 0.29 | 95\% |
| 2 | rs4851004 | 103,009,537 | C/T | 0.49 | 0.92 (0.89-0.94) | $4.45 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.82 \times 10^{-11}$ | 0.29 | 100\% |
| 2 | rs2287034 | 103,010,588 | C/A | 0.29 | 1.11 (1.08-1.14) | $3.58 \times 10^{-14}$ | 1.11 (1.08-1.14) | $3.58 \times 10^{-14}$ | 0.61 | 100\% |
| 2 | rs2287033 | 103,011,237 | T/C | 0.49 | 0.92 (0.89-0.94) | $4.12 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.78 \times 10^{-11}$ | 0.29 | 100\% |
| 2 | rs4851005 | 103,011,552 | C/T | 0.38 | 1.09 (1.06-1.12) | $5.40 \times 10^{-9}$ | 1.09 (1.06-1.12) | $8.91 \times 10^{-11}$ | 0.26 | 100\% |
| 2 | rs1135354 | 103,014,302 | T/G | 0.29 | 1.11 (1.08-1.14) | $6.31 \times 10^{-13}$ | 1.11 (1.08-1.14) | $6.31 \times 10^{-13}$ | 0.52 | 91\% |
| 2 | rs1420094 | 103,015,687 | C/T | 0.49 | 0.92 (0.89-0.94) | $3.82 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.54 \times 10^{-11}$ | 0.29 | 100\% |
| 2 | rs17027087 | 103,015,918 | C/T | 0.29 | 1.11 (1.08-1.14) | $4.18 \times 10^{-14}$ | 1.11 (1.08-1.14) | $4.18 \times 10^{-14}$ | 0.57 | 100\% |
| 2 | rs6710528 | 103,016,142 | C/T | 0.49 | 0.92 (0.89-0.94) | $9.76 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.83 \times 10^{-11}$ | 0.23 | 100\% |
| 2 | rs3732124 | 103,018,052 | C/T | 0.49 | 0.92 (0.89-0.94) | $7.18 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.55 \times 10^{-11}$ | 0.24 | 100\% |
| 2 | rs4851571 | 103,019,000 | C/T | 0.49 | 0.92 (0.89-0.94) | $7.45 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.74 \times 10^{-11}$ | 0.25 | 100\% |
| 2 | rs4851572 | 103,019,031 | G/A | 0.49 | 0.92 (0.89-0.94) | $7.46 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.72 \times 10^{-11}$ | 0.25 | 100\% |
| 2 | rs6710034 | 103,023,678 | G/A | 0.49 | 0.92 (0.89-0.94) | $3.29 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.32 \times 10^{-11}$ | 0.29 | 100\% |
| 2 | rs10203558 | 103,027,640 | T/C | 0.49 | 0.92 (0.89-0.94) | $3.27 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.31 \times 10^{-11}$ | 0.29 | 100\% |
| 2 | rs10200952 | 103,027,651 | A/C | 0.49 | 0.91 (0.89-0.94) | $1.77 \times 10^{-10}$ | 0.92 (0.89-0.94) | $2.53 \times 10^{-11}$ | 0.37 | 91\% |
| 2 | rs1420105 | 103,035,119 | T/C | 0.49 | 0.92 (0.89-0.94) | $2.25 \times 10^{-10}$ | 0.92 (0.9-0.94) | $7.94 \times 10^{-12}$ | 0.28 | 100\% |
| 2 | rs2293224 | 103,035,779 | T/C | 0.49 | 0.92 (0.89-0.94) | $4.45 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.32 \times 10^{-11}$ | 0.27 | 100\% |
| 2 | rs6743516 | 103,036,335 | A/G | 0.49 | 0.92 (0.89-0.94) | $4.31 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.36 \times 10^{-11}$ | 0.27 | 100\% |
| 2 | rs3771156 | 103,036,677 | C/T | 0.29 | 1.11 (1.08-1.14) | $2.67 \times 10^{-14}$ | 1.11 (1.08-1.14) | $2.67 \times 10^{-14}$ | 0.53 | 100\% |
| 2 | rs1420100 | 103,037,002 | C/A | 0.49 | 0.92 (0.89-0.94) | $4.22 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.50 \times 10^{-11}$ | 0.28 | 100\% |
| 2 | rs3771155 | 103,037,826 | A/G | 0.49 | 0.92 (0.89-0.94) | $3.77 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.06 \times 10^{-11}$ | 0.27 | 100\% |
| 2 | rs10206291 | 103,038,863 | T/C | 0.49 | 0.92 (0.89-0.94) | $4.23 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.44 \times 10^{-11}$ | 0.28 | 100\% |
| 2 | rs885088 | 103,039,044 | A/G | 0.49 | 0.92 (0.89-0.94) | $3.95 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.36 \times 10^{-11}$ | 0.28 | 100\% |
| 2 | rs3771154 | 103,039,360 | C/T | 0.49 | 0.92 (0.89-0.94) | $4.33 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.36 \times 10^{-11}$ | 0.27 | 100\% |
| 2 | rs6759479 | 103,040,047 | A/C | 0.49 | 0.92 (0.89-0.94) | $5.05 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.93 \times 10^{-11}$ | 0.27 | 100\% |
| 2 | rs887972 | 103,040,945 | G/A | 0.30 | 1.11 (1.08-1.14) | $4.25 \times 10^{-12}$ | 1.11 (1.08-1.14) | $4.25 \times 10^{-12}$ | 0.60 | 89\% |
| 2 | rs887971 | 103,041,167 | T/C | 0.30 | 1.11 (1.08-1.14) | $3.79 \times 10^{-14}$ | 1.11 (1.08-1.14) | $3.79 \times 10^{-14}$ | 0.60 | 100\% |
| 2 | rs3755266 | 103,042,712 | G/A | 0.49 | 0.92 (0.89-0.94) | $2.05 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.49 \times 10^{-11}$ | 0.33 | 100\% |
| 2 | rs7559845 | 103,046,214 | T/G | 0.49 | 0.92 (0.89-0.94) | $3.01 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.54 \times 10^{-11}$ | 0.31 | 100\% |
| 2 | rs2310300 | 103,049,074 | A/G | 0.49 | 0.92 (0.89-0.94) | $1.10 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.09 \times 10^{-11}$ | 0.35 | 100\% |
| 2 | rs11681718 | 103,051,144 | A/G | 0.29 | 1.1 (1.07-1.13) | $6.61 \times 10^{-13}$ | 1.1 (1.07-1.13) | $4.62 \times 10^{-13}$ | 0.46 | 100\% |
| 2 | rs4851582 | 103,051,558 | T/C | 0.29 | 1.1 (1.07-1.13) | $6.39 \times 10^{-13}$ | 1.1 (1.07-1.13) | $4.56 \times 10^{-13}$ | 0.46 | 100\% |
| 2 | rs3755265 | 103,052,816 | C/A | 0.49 | 0.92 (0.89-0.94) | $4.67 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.90 \times 10^{-11}$ | 0.29 | 100\% |
| 2 | rs2058659 | 103,054,556 | G/A | 0.49 | 0.92 (0.9-0.94) | $1.64 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.93 \times 10^{-11}$ | 0.36 | 100\% |
| 2 | rs17027166 | 103,055,420 | G/A | 0.29 | 1.1 (1.07-1.13) | $5.74 \times 10^{-13}$ | 1.1 (1.07-1.13) | $5.74 \times 10^{-13}$ | 0.48 | 100\% |
| 2 | rs13021177 | 103,056,493 | A/G | 0.49 | 0.92 (0.9-0.94) | $4.09 \times 10^{-10}$ | 0.92 (0.9-0.94) | $2.26 \times 10^{-11}$ | 0.31 | 100\% |
| 2 | rs10490204 | 103,056,534 | A/C | 0.29 | 1.1 (1.08-1.13) | $2.35 \times 10^{-13}$ | 1.1 (1.08-1.13) | $2.35 \times 10^{-13}$ | 0.50 | 100\% |
| 2 | rs17027179 | 103,057,159 | C/T | 0.29 | 1.1 (1.07-1.13) | $3.90 \times 10^{-13}$ | 1.1 (1.07-1.13) | $3.90 \times 10^{-13}$ | 0.48 | 100\% |
| 2 | rs10490203 | 103,059,237 | T/G | 0.29 | 1.1 (1.07-1.13) | $3.57 \times 10^{-13}$ | 1.1 (1.07-1.13) | $3.57 \times 10^{-13}$ | 0.48 | 100\% |
| 2 | rs3771150 | 103,060,851 | G/A | 0.29 | 1.1 (1.07-1.13) | $3.93 \times 10^{-13}$ | 1.1 (1.07-1.13) | $3.93 \times 10^{-13}$ | 0.49 | 100\% |
| 2 | rs11123928 | 103,061,286 | G/A | 0.34 | 1.08 (1.06-1.12) | $6.75 \times 10^{-9}$ | 1.08 (1.06-1.12) | $6.75 \times 10{ }^{-9}$ | 0.51 | 96\% |
| 2 | rs7597017 | 103,062,116 | A/G | 0.34 | 1.08 (1.06-1.12) | $6.54 \times 10^{-9}$ | 1.08 (1.06-1.12) | $6.54 \times 10^{-9}$ | 0.51 | 96\% |
| 2 | rs17027230 | 103,079,330 | C/T | 0.29 | 1.1 (1.07-1.13) | $2.36 \times 10^{-12}$ | 1.1 (1.07-1.13) | $7.82 \times 10^{-13}$ | 0.43 | 98\% |


| Chrom | SNP | Position | Allele (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 | rs4851011 | 103,089,678 | C/T | 0.29 | 1.1 (1.07-1.13) | $1.02 \times 10^{-11}$ | 1.1 (1.07-1.13) | $1.59 \times 10^{-12}$ | 0.40 | 98\% |
| 2 | rs17027255 | 103,090,127 | C/T | 0.29 | 1.1 (1.07-1.13) | $6.90 \times 10^{-12}$ | 1.1 (1.07-1.13) | $1.61 \times 10^{-12}$ | 0.42 | 98\% |
| 2 | rs17027258 | 103,091,540 | A/G | 0.29 | 1.1 (1.07-1.13) | $7.72 \times 10^{-12}$ | 1.1 (1.07-1.13) | $1.60 \times 10^{-12}$ | 0.41 | 98\% |
| 5 | rs17446534 | 110,079,450 | T/C | 0.09 | 1.14 (1.09-1.18) | $4.02 \times 10^{-9}$ | 1.14 (1.09-1.18) | $4.02 \times 10^{-9}$ | 0.82 | 100\% |
| 5 | rs244923 | 110,080,363 | T/C | 0.91 | 0.88 (0.84-0.92) | $6.96 \times 10^{-9}$ | 0.88 (0.84-0.92) | $6.96 \times 10^{-9}$ | 0.79 | 98\% |
| 5 | rs381661 | 110,085,098 | C/T | 0.90 | 0.86 (0.82-0.9) | $2.19 \times 10^{-10}$ | 0.86 (0.82-0.9) | $2.19 \times 10^{-10}$ | 0.88 | 96\% |
| 5 | rs6893213 | 110,198,114 | C/T | 0.08 | 1.15 (1.1-1.21) | $6.22 \times 10^{-9}$ | 1.16 (1.11-1.21) | $7.99 \times 10^{-11}$ | 0.28 | 98\% |
| 5 | rs1837253 | 110,401,872 | T/C | 0.74 | 1.16 (1.12-1.21) | $4.94 \times 10^{-15}$ | 1.16 (1.12-1.19) | $1.06 \times 10^{-21}$ | 0.06 | 96\% |
| 5 | rs10455025 | 110,404,999 | A/C | 0.34 | 1.15 (1.12-1.18) | $2.03 \times 10^{-25}$ | 1.15 (1.12-1.18) | $2.03 \times 10^{-25}$ | 0.53 | 98\% |
| 5 | rs3806932 | 110,405,675 | A/G | 0.45 | 0.92 (0.9-0.95) | $4.51 \times 10^{-10}$ | 0.92 (0.9-0.95) | $2.49 \times 10^{-10}$ | 0.45 | 100\% |
| 5 | rs3806933 | 110,406,742 | C/T | 0.45 | 0.92 (0.9-0.95) | $2.70 \times 10^{-10}$ | 0.92 (0.9-0.95) | $1.42 \times 10^{-10}$ | 0.45 | 100\% |
| 5 | rs1898671 | 110,408,002 | C/T | 0.34 | 1.15 (1.12-1.18) | $7.12 \times 10^{-25}$ | 1.15 (1.12-1.18) | $7.12 \times 10^{-25}$ | 0.50 | 100\% |
| 5 | rs7723819 | 110,427,347 | G/A | 0.47 | 0.92 (0.89-0.94) | $1.00 \times 10^{-9}$ | 0.92 (0.89-0.94) | $3.05 \times 10^{-12}$ | 0.26 | 100\% |
| 5 | rs1993465 | 110,433,098 | A/G | 0.46 | 0.92 (0.9-0.94) | $2.32 \times 10^{-10}$ | 0.92 (0.9-0.94) | $1.10 \times 10^{-11}$ | 0.36 | 100\% |
| 5 | rs6859041 | 110,435,231 | G/A | 0.47 | 0.92 (0.89-0.94) | $1.02 \times 10^{-10}$ | 0.92 (0.89-0.94) | $4.71 \times 10^{-12}$ | 0.36 | 100\% |
| 5 | rs10038177 | 110,436,450 | C/T | 0.47 | 0.92 (0.89-0.94) | $1.37 \times 10^{-10}$ | 0.92 (0.89-0.94) | $4.63 \times 10^{-12}$ | 0.35 | 100\% |
| 5 | rs10045255 | 110,438,357 | A/G | 0.47 | 0.92 (0.89-0.94) | $2.03 \times 10^{-10}$ | 0.92 (0.89-0.94) | $2.85 \times 10^{-12}$ | 0.32 | 100\% |
| 5 | rs1379300 | 110,441,439 | T/C | 0.47 | 0.92 (0.89-0.94) | $5.58 \times 10^{-10}$ | 0.92 (0.89-0.94) | $4.96 \times 10^{-12}$ | 0.29 | 100\% |
| 5 | rs10038058 | 110,443,281 | A/G | 0.47 | 0.92 (0.89-0.94) | $7.49 \times 10^{-10}$ | 0.92 (0.89-0.94) | $4.20 \times 10^{-12}$ | 0.27 | 100\% |
| 5 | rs2112541 | 110,449,346 | T/C | 0.47 | 0.92 (0.89-0.94) | $1.92 \times 10^{-9}$ | 0.92 (0.9-0.94) | $5.00 \times 10^{-12}$ | 0.23 | 100\% |
| 5 | rs10060003 | 110,449,357 | A/G | 0.47 | 0.92 (0.89-0.95) | $2.50 \times 10^{-9}$ | 0.92 (0.9-0.94) | $5.54 \times 10^{-12}$ | 0.23 | 100\% |
| 5 | rs10051830 | 110,452,845 | G/A | 0.46 | 0.92 (0.9-0.95) | $6.16 \times 10^{-9}$ | 0.92 (0.9-0.94) | $2.72 \times 10^{-11}$ | 0.23 | 100\% |
| 5 | rs17624673 | 110,457,158 | C/T | 0.33 | 1.14 (1.11-1.17) | $1.51 \times 10^{-19}$ | 1.14 (1.11-1.17) | $9.65 \times 10^{-23}$ | 0.32 | 98\% |
| 5 | rs7702774 | 110,460,851 | G/T | 0.47 | 0.92 (0.89-0.94) | $1.43 \times 10^{-9}$ | 0.92 (0.9-0.94) | $6.66 \times 10^{-12}$ | 0.26 | 100\% |
| 5 | rs4530809 | 110,462,230 | A/G | 0.37 | 0.93 (0.9-0.95) | $5.53 \times 10^{-9}$ | 0.93 (0.9-0.95) | $5.53 \times 10^{-9}$ | 0.84 | 98\% |
| 5 | rs1043828 | 110,464,008 | T/C | 0.34 | 1.13 (1.1-1.17) | $3.01 \times 10^{-19}$ | 1.13 (1.11-1.16) | $1.48 \times 10^{-22}$ | 0.31 | 100\% |
| 5 | rs1438673 | 110,467,499 | C/T | 0.51 | 0.91 (0.88-0.93) | $4.52 \times 10^{-11}$ | 0.9 (0.88-0.92) | $2.50 \times 10^{-16}$ | 0.12 | 100\% |
| 5 | rs6594499 | 110,470,137 | C/A | 0.51 | 0.9 (0.88-0.93) | $3.84 \times 10^{-11}$ | 0.9 (0.88-0.92) | $6.30 \times 10^{-16}$ | 0.15 | 96\% |
| 5 | rs2162893 | 110,558,064 | A/G | 0.62 | 0.9 (0.88-0.93) | $3.95 \times 10^{-13}$ | 0.9 (0.88-0.93) | $3.95 \times 10^{-13}$ | 0.64 | 91\% |
| 5 | rs10057913 | 110,566,175 | G/A | 0.88 | 1.13 (1.08-1.17) | $2.00 \times 10^{-9}$ | 1.13 (1.08-1.17) | $2.00 \times 10^{-9}$ | 0.77 | 98\% |
| 5 | rs1457115 | 110,567,598 | C/T | 0.45 | 1.08 (1.06-1.11) | $1.78 \times 10^{-10}$ | 1.08 (1.06-1.11) | $1.78 \times 10^{-10}$ | 0.94 | 98\% |
| 5 | rs11950562 | 131,652,529 | A/C | 0.45 | 0.93 (0.9-0.95) | $1.65 \times 10^{-9}$ | 0.93 (0.9-0.95) | $1.65 \times 10^{-9}$ | 0.63 | 100\% |
| 5 | rs10058074 | 131,686,146 | G/A | 0.45 | 0.93 (0.91-0.95) | $3.89 \times 10^{-9}$ | 0.93 (0.91-0.95) | $3.89 \times 10^{-9}$ | 0.53 | 100\% |
| 5 | rs4705938 | 131,694,077 | T/C | 0.45 | 0.93 (0.9-0.95) | $3.97 \times 10^{-9}$ | 0.93 (0.9-0.95) | $3.25 \times 10^{-9}$ | 0.46 | 100\% |
| 5 | rs17622208 | 131,717,050 | G/A | 0.45 | 0.93 (0.9-0.95) | $1.02 \times 10^{-9}$ | 0.93 (0.9-0.95) | $1.02 \times 10^{-9}$ | 0.51 | 100\% |
| 5 | rs1023518 | 131,793,772 | G/T | 0.27 | 1.09 (1.06-1.12) | $1.16 \times 10^{-9}$ | 1.09 (1.06-1.12) | $1.16 \times 10^{-9}$ | 0.59 | 100\% |
| 5 | rs3857440 | 131,794,069 | G/A | 0.27 | 1.09 (1.06-1.12) | $1.61 \times 10^{-9}$ | 1.09 (1.06-1.12) | $1.61 \times 10^{-9}$ | 0.57 | 100\% |
| 5 | rs11745587 | 131,796,922 | G/A | 0.36 | 1.09 (1.06-1.11) | $3.19 \times 10^{-10}$ | 1.09 (1.06-1.11) | $3.19 \times 10^{-10}$ | 0.54 | 98\% |
| 5 | rs6894249 | 131,797,547 | A/G | 0.39 | 1.09 (1.06-1.12) | $2.15 \times 10^{-11}$ | 1.09 (1.06-1.12) | $2.15 \times 10^{-11}$ | 0.48 | 96\% |
| 5 | rs17622656 | 131,820,997 | G/A | 0.38 | 0.91 (0.88-0.94) | $1.00 \times 10^{-9}$ | 0.91 (0.89-0.94) | $1.94 \times 10^{-11}$ | 0.26 | 96\% |
| 5 | rs736801 | 131,833,599 | C/T | 0.38 | 0.91 (0.88-0.94) | $2.86 \times 10^{-10}$ | 0.91 (0.89-0.94) | $8.51 \times 10^{-12}$ | 0.29 | 96\% |
| 5 | rs2158177 | 131,984,058 | A/G | 0.18 | 1.13 (1.09-1.18) | $2.36 \times 10^{-9}$ | 1.13 (1.09-1.17) | $6.18 \times 10^{-11}$ | 0.26 | 96\% |
| 5 | rs1295686 | 131,995,843 | T/C | 0.78 | 0.89 (0.87-0.92) | $4.11 \times 10^{-13}$ | 0.89 (0.87-0.92) | $4.11 \times 10^{-13}$ | 0.57 | 100\% |
| 5 | rs20541 | 131,995,964 | A/G | 0.79 | 0.89 (0.86-0.91) | $1.36 \times 10^{-14}$ | 0.89 (0.86-0.91) | $1.36 \times 10^{-14}$ | 0.73 | 98\% |
| 5 | rs1295685 | 131,996,445 | A/G | 0.78 | 0.89 (0.86-0.92) | $2.24 \times 10^{-13}$ | 0.89 (0.86-0.92) | $1.21 \times 10^{-13}$ | 0.45 | 98\% |
| 5 | rs848 | 131,996,500 | A/C | 0.79 | 0.89 (0.86-0.92) | $1.01 \times 10^{-12}$ | 0.89 (0.86-0.92) | $2.60 \times 10^{-13}$ | 0.41 | 96\% |
| 5 | rs847 | 131,996,669 | T/C | 0.79 | 0.89 (0.86-0.92) | $4.66 \times 10^{-12}$ | 0.89 (0.86-0.92) | $8.53 \times 10^{-13}$ | 0.38 | 98\% |
| 6 | rs16901847 | 28,375,851 | G/A | 0.19 | 1.1 (1.06-1.13) | $9.42 \times 10^{-9}$ | 1.1 (1.06-1.13) | $9.42 \times 10^{-9}$ | 0.79 | 100\% |
| 6 | rs7765989 | 28,400,295 | T/C | 0.18 | 1.1 (1.06-1.13) | $9.61 \times 10^{-9}$ | 1.1 (1.06-1.13) | $9.61 \times 10^{-9}$ | 0.78 | 100\% |
| 6 | rs16894116 | 28,414,967 | G/T | 0.18 | 1.1 (1.06-1.13) | $9.27 \times 10^{-9}$ | 1.1 (1.06-1.13) | $9.27 \times 10^{-9}$ | 0.75 | 100\% |
| 6 | rs1233578 | 28,712,247 | A/G | 0.13 | 1.11 (1.07-1.15) | $5.32 \times 10^{-9}$ | 1.11 (1.07-1.15) | $5.32 \times 10^{-9}$ | 0.82 | 100\% |
| 6 | rs3130117 | 30,508,956 | G/A | 0.11 | 1.12 (1.08-1.17) | $9.09 \times 10^{-9}$ | 1.12 (1.08-1.17) | $9.09 \times 10^{-9}$ | 0.59 | 100\% |
| 6 | rs3132610 | 30,544,401 | A/G | 0.11 | 1.12 (1.08-1.17) | $8.07 \times 10^{-9}$ | 1.12 (1.08-1.17) | $8.07 \times 10^{-9}$ | 0.65 | 100\% |
| 6 | rs9262143 | 30,652,781 | C/T | 0.11 | 1.12 (1.08-1.17) | $5.97 \times 10^{-9}$ | 1.12 (1.08-1.17) | $5.97 \times 10^{-9}$ | 0.67 | 100\% |
| 6 | rs3132584 | 30,688,427 | G/T | 0.18 | 1.1 (1.07-1.14) | $4.06 \times 10^{-10}$ | 1.1 (1.07-1.14) | $4.06 \times 10^{-10}$ | 0.77 | 100\% |
| 6 | rs8233 | 30,692,965 | G/A | 0.18 | 1.11 (1.07-1.14) | $3.03 \times 10^{-10}$ | 1.11 (1.07-1.14) | $3.03 \times 10^{-10}$ | 0.76 | 100\% |
| 6 | rs3095330 | 30,693,633 | A/G | 0.17 | 1.12 (1.08-1.15) | $5.73 \times 10^{-10}$ | 1.12 (1.08-1.15) | $5.73 \times 10^{-10}$ | 0.78 | 89\% |
| 6 | rs3095329 | 30,693,816 | A/G | 0.18 | 1.11 (1.07-1.14) | $3.07 \times 10^{-10}$ | 1.11 (1.07-1.14) | $3.07 \times 10^{-10}$ | 0.77 | 100\% |
| 6 | rs3094127 | 30,697,447 | A/G | 0.18 | 1.11 (1.07-1.14) | $2.67 \times 10^{-10}$ | 1.11 (1.07-1.14) | $2.67 \times 10^{-10}$ | 0.77 | 100\% |
| 6 | rs1064627 | 30,698,541 | A/G | 0.18 | 1.11 (1.07-1.14) | $4.59 \times 10^{-10}$ | 1.11 (1.07-1.14) | $4.59 \times 10^{-10}$ | 0.74 | 98\% |


| Chrom | SNP | Position | Allele (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 6 | rs1059612 | 30,708,955 | C/T | 0.12 | 1.13 (1.08-1.17) | $1.56 \times 10^{-9}$ | 1.13 (1.08-1.17) | $1.56 \times 10-9$ | 0.60 | 100\% |
| 6 | rs2284174 | 30,713,580 | T/C | 0.16 | 1.11 (1.07-1.15) | $9.37 \times 10^{-10}$ | 1.11 (1.07-1.15) | $9.37 \times 10^{-10}$ | 0.68 | 100\% |
| 6 | rs3129973 | 30,721,143 | C/T | 0.11 | 1.12 (1.08-1.16) | $8.26 \times 10^{-9}$ | 1.12 (1.08-1.16) | $8.26 \times 10^{-9}$ | 0.63 | 100\% |
| 6 | rs3130665 | 30,735,979 | C/T | 0.12 | 1.12 (1.08-1.16) | $4.67 \times 10^{-9}$ | 1.12 (1.08-1.16) | $4.67 \times 10^{-9}$ | 0.56 | 100\% |
| 6 | rs3095336 | 30,738,446 | G/A | 0.12 | 1.12 (1.08-1.17) | $2.87 \times 10^{-9}$ | 1.12 (1.08-1.17) | $2.87 \times 10^{-9}$ | 0.77 | 98\% |
| 6 | rs3132605 | 30,739,972 | A/C | 0.12 | 1.12 (1.08-1.16) | $3.25 \times 10^{-9}$ | 1.12 (1.08-1.16) | $3.25 \times 10^{-9}$ | 0.76 | 98\% |
| 6 | rs3129978 | 30,746,331 | A/G | 0.12 | 1.12 (1.08-1.17) | $3.11 \times 10^{-9}$ | 1.12 (1.08-1.17) | $3.11 \times 10^{-9}$ | 0.75 | 98\% |
| 6 | rs3130673 | 30,746,519 | G/T | 0.12 | 1.12 (1.08-1.17) | $2.56 \times 10^{-9}$ | 1.12 (1.08-1.17) | $2.56 \times 10^{-9}$ | 0.75 | 98\% |
| 6 | rs3131044 | 30,758,664 | A/C | 0.12 | 1.12 (1.08-1.17) | $2.17 \times 10^{-9}$ | 1.12 (1.08-1.17) | $2.17 \times 10^{-9}$ | 0.71 | 98\% |
| 6 | rs3131050 | 30,760,025 | T/C | 0.12 | 1.12 (1.08-1.17) | $3.71 \times 10^{-9}$ | 1.12 (1.08-1.17) | $3.71 \times 10^{-9}$ | 0.70 | 96\% |
| 6 | rs9262200 | 30,760,725 | G/A | 0.12 | 1.12 (1.08-1.17) | $2.62 \times 10^{-9}$ | 1.12 (1.08-1.17) | $2.62 \times 10^{-9}$ | 0.73 | 100\% |
| 6 | rs3131055 | 30,761,487 | T/C | 0.12 | 1.12 (1.08-1.16) | $2.98 \times 10^{-9}$ | 1.12 (1.08-1.16) | $2.98 \times 10^{-9}$ | 0.72 | 100\% |
| 6 | rs3129985 | 30,762,542 | C/T | 0.11 | 1.12 (1.08-1.16) | $3.21 \times 10^{-9}$ | 1.12 (1.08-1.16) | $3.21 \times 10^{-9}$ | 0.72 | 100\% |
| 6 | rs3131060 | 30,763,291 | G/A | 0.11 | 1.12 (1.08-1.16) | $2.96 \times 10^{-9}$ | 1.12 (1.08-1.16) | $2.96 \times 10^{-9}$ | 0.72 | 100\% |
| 6 | rs3131064 | 30,763,893 | T/C | 0.14 | 1.12 (1.08-1.16) | $2.58 \times 10^{-10}$ | 1.12 (1.08-1.16) | $2.58 \times 10^{-10}$ | 0.54 | 100\% |
| 6 | rs3130641 | 30,764,081 | C/T | 0.11 | 1.12 (1.08-1.17) | $6.59 \times 10^{-9}$ | 1.12 (1.08-1.17) | $6.59 \times 10^{-9}$ | 0.74 | 96\% |
| 6 | rs1264376 | 30,765,579 | C/A | 0.12 | 1.12 (1.08-1.16) | $4.62 \times 10^{-9}$ | 1.12 (1.08-1.16) | $4.62 \times 10^{-9}$ | 0.69 | 98\% |
| 6 | rs3130564 | 31,101,674 | C/T | 0.19 | 1.1 (1.06-1.13) | $3.71 \times 10^{-9}$ | 1.1 (1.06-1.13) | $3.71 \times 10^{-9}$ | 0.68 | 98\% |
| 6 | rs9264740 | 31,244,331 | C/T | 0.72 | 1.09 (1.06-1.12) | $9.25 \times 10^{-9}$ | 1.09 (1.06-1.12) | $9.25 \times 10^{-9}$ | 0.68 | 89\% |
| 6 | rs2596464 | 31,412,961 | T/C | 0.48 | 1.1 (1.06-1.13) | $1.33 \times 10^{-9}$ | 1.1 (1.07-1.13) | $1.79 \times 10^{-13}$ | 0.14 | 98\% |
| 6 | rs2596472 | 31,428,967 | G/A | 0.73 | 1.1 (1.07-1.13) | $5.54 \times 10^{-10}$ | 1.1 (1.07-1.13) | $5.54 \times 10^{-10}$ | 0.79 | 95\% |
| 6 | rs1055388 | 31,501,737 | C/T | 0.72 | 1.09 (1.06-1.12) | $4.98 \times 10^{-9}$ | 1.09 (1.06-1.12) | $4.98 \times 10^{-9}$ | 0.83 | 100\% |
| 6 | rs933208 | 31,506,648 | G/T | 0.72 | 1.09 (1.06-1.12) | $4.16 \times 10^{-9}$ | 1.09 (1.06-1.12) | $4.16 \times 10^{-9}$ | 0.83 | 100\% |
| 6 | rs2239528 | 31,510,105 | A/G | 0.72 | 1.09 (1.06-1.12) | $6.14 \times 10^{-9}$ | 1.09 (1.06-1.12) | $6.14 \times 10^{-9}$ | 0.84 | 100\% |
| 6 | rs2523504 | 31,510,858 | T/C | 0.72 | 1.09 (1.06-1.12) | $6.05 \times 10^{-9}$ | 1.09 (1.06-1.12) | $6.05 \times 10^{-9}$ | 0.84 | 100\% |
| 6 | rs6904608 | 32,327,727 | C/T | 0.56 | 1.08 (1.05-1.1) | $4.84 \times 10^{-9}$ | 1.08 (1.05-1.1) | $4.84 \times 10^{-9}$ | 0.84 | 100\% |
| 6 | rs2073044 | 32,338,986 | C/T | 0.24 | 1.11 (1.08-1.15) | $1.24 \times 10^{-10}$ | 1.12 (1.08-1.15) | $4.31 \times 10^{-13}$ | 0.30 | 93\% |
| 6 | rs9268403 | 32,341,473 | T/C | 0.27 | 1.13 (1.09-1.16) | $4.95 \times 10^{-13}$ | 1.13 (1.1-1.16) | $5.79 \times 10^{-18}$ | 0.14 | 98\% |
| 6 | rs9268429 | 32,345,052 | A/G | 0.27 | 1.12 (1.09-1.16) | $1.35 \times 10^{-12}$ | 1.13 (1.09-1.16) | $3.03 \times 10^{-17}$ | 0.16 | 98\% |
| 6 | rs1980495 | 32,346,794 | A/C | 0.26 | 1.13 (1.09-1.17) | $3.55 \times 10^{-11}$ | 1.13 (1.1-1.17) | $1.76 \times 10^{-16}$ | 0.11 | 89\% |
| 6 | rs8180664 | 32,347,490 | C/T | 0.26 | 1.13 (1.09-1.17) | $1.10 \times 10^{-13}$ | 1.13 (1.1-1.16) | $4.39 \times 10^{-19}$ | 0.13 | 100\% |
| 6 | rs2395157 | 32,348,145 | A/G | 0.26 | 1.13 (1.1-1.17) | $9.64 \times 10^{-14}$ | 1.13 (1.1-1.16) | $1.92 \times 10^{-19}$ | 0.11 | 100\% |
| 6 | rs17423649 | 32,357,133 | C/T | 0.16 | 0.89 (0.86-0.92) | $2.68 \times 10^{-10}$ | 0.89 (0.86-0.92) | $2.68 \times 10^{-10}$ | 0.49 | 98\% |
| 6 | rs9268474 | 32,357,165 | T/C | 0.27 | 1.13 (1.09-1.17) | $6.29 \times 10^{-13}$ | 1.13 (1.1-1.16) | $8.10 \times 10^{-19}$ | 0.09 | 100\% |
| 6 | rs12529049 | 32,357,715 | C/T | 0.16 | 0.89 (0.86-0.92) | $1.64 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.64 \times 10^{-10}$ | 0.50 | 100\% |
| 6 | rs12525722 | 32,358,163 | A/G | 0.16 | 0.89 (0.86-0.92) | $1.79 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.79 \times 10^{-10}$ | 0.51 | 100\% |
| 6 | rs17423691 | 32,358,345 | C/A | 0.16 | 0.89 (0.86-0.92) | $1.73 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.73 \times 10^{-10}$ | 0.51 | 100\% |
| 6 | rs17423698 | 32,358,368 | T/C | 0.16 | 0.89 (0.86-0.92) | $1.68 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.68 \times 10^{-10}$ | 0.51 | 100\% |
| 6 | rs17495592 | 32,358,533 | A/G | 0.16 | 0.89 (0.86-0.92) | $1.51 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.51 \times 10^{-10}$ | 0.51 | 100\% |
| 6 | rs9268477 | 32,359,121 | G/A | 0.27 | 1.12 (1.09-1.16) | $5.65 \times 10^{-11}$ | 1.13 (1.09-1.16) | $3.11 \times 10^{-16}$ | 0.10 | 91\% |
| 6 | rs17495612 | 32,359,431 | T/C | 0.16 | 0.89 (0.86-0.92) | $1.11 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.11 \times 10^{-10}$ | 0.56 | 100\% |
| 6 | rs16870123 | 32,359,460 | G/A | 0.16 | 0.89 (0.86-0.92) | $1.27 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.27 \times 10^{-10}$ | 0.54 | 100\% |
| 6 | rs17423753 | 32,360,341 | C/A | 0.16 | 0.89 (0.86-0.93) | $3.41 \times 10^{-10}$ | 0.89 (0.86-0.93) | $3.41 \times 10^{-10}$ | 0.53 | 100\% |
| 6 | rs3817969 | 32,361,388 | C/T | 0.16 | 0.89 (0.86-0.92) | $2.16 \times 10^{-10}$ | 0.89 (0.86-0.92) | $2.16 \times 10^{-10}$ | 0.48 | 96\% |
| 6 | rs9268480 | 32,363,844 | C/T | 0.27 | 1.13 (1.09-1.17) | $5.88 \times 10^{-13}$ | 1.13 (1.1-1.16) | $7.24 \times 10^{-19}$ | 0.09 | 100\% |
| 6 | rs4248166 | 32,366,421 | T/C | 0.20 | 0.9 (0.87-0.93) | $2.65 \times 10^{-10}$ | 0.9 (0.87-0.93) | $2.65 \times 10^{-10}$ | 0.50 | 100\% |
| 6 | rs2294881 | 32,367,604 | T/C | 0.20 | 0.91 (0.88-0.93) | $8.49 \times 10^{-10}$ | 0.91 (0.88-0.93) | $8.49 \times 10^{-10}$ | 0.59 | 100\% |
| 6 | rs2294880 | 32,367,722 | A/G | 0.26 | 1.13 (1.09-1.17) | $4.81 \times 10^{-10}$ | 1.13 (1.1-1.17) | $1.24 \times 10^{-15}$ | 0.09 | 86\% |
| 6 | rs3817966 | 32,367,847 | T/C | 0.27 | 1.13 (1.08-1.17) | $4.80 \times 10^{-10}$ | 1.13 (1.1-1.16) | $6.29 \times 10^{-16}$ | 0.06 | 89\% |
| 6 | rs3817963 | 32,368,087 | T/C | 0.27 | 1.13 (1.09-1.17) | $7.76 \times 10^{-13}$ | 1.13 (1.1-1.16) | $8.10 \times 10^{-19}$ | 0.08 | 100\% |
| 6 | rs3817962 | 32,368,314 | C/A | 0.26 | 1.13 (1.08-1.17) | $6.80 \times 10^{-10}$ | 1.13 (1.1-1.17) | $1.71 \times 10^{-15}$ | 0.08 | 86\% |
| 6 | rs2076525 | 32,370,616 | T/C | 0.27 | 1.12 (1.09-1.16) | $1.08 \times 10^{-11}$ | 1.13 (1.1-1.16) | $2.91 \times 10^{-17}$ | 0.09 | 96\% |
| 6 | rs2076524 | 32,370,684 | A/G | 0.26 | 1.13 (1.09-1.17) | $1.33 \times 10^{-13}$ | 1.13 (1.1-1.16) | $2.80 \times 10^{-19}$ | 0.11 | 100\% |
| 6 | rs3793126 | 32,371,619 | A/G | 0.28 | 1.13 (1.09-1.16) | $5.78 \times 10^{-13}$ | 1.13 (1.1-1.16) | $2.31 \times 10^{-18}$ | 0.09 | 100\% |
| 6 | rs3793127 | 32,371,915 | C/T | 0.21 | 1.13 (1.09-1.17) | $1.74 \times 10^{-11}$ | 1.14 (1.11-1.17) | $4.88 \times 10^{-18}$ | 0.09 | 100\% |
| 6 | rs9268493 | 32,375,330 | G/A | 0.28 | 1.13 (1.09-1.16) | $2.12 \times 10^{-11}$ | 1.13 (1.09-1.16) | $7.47 \times 10^{-17}$ | 0.07 | 95\% |
| 6 | rs9268494 | 32,375,352 | A/C | 0.27 | 1.13 (1.09-1.16) | $5.70 \times 10^{-12}$ | 1.13 (1.1-1.16) | $4.96 \times 10^{-17}$ | 0.11 | 96\% |
| 6 | rs9268497 | 32,375,424 | G/A | 0.27 | 1.13 (1.09-1.17) | $7.77 \times 10^{-11}$ | 1.13 (1.1-1.17) | $1.54 \times 10^{-16}$ | 0.09 | 89\% |
| 6 | rs9268499 | 32,375,695 | G/A | 0.28 | 1.13 (1.09-1.17) | $1.41 \times 10^{-11}$ | 1.12 (1.09-1.16) | $9.06 \times 10^{-17}$ | 0.07 | 95\% |
| 6 | rs3763309 | 32,375,973 | C/A | 0.21 | 1.13 (1.09-1.18) | $3.31 \times 10^{-12}$ | 1.14 (1.11-1.18) | $1.31 \times 10^{-18}$ | 0.11 | 100\% |


| Chrom | SNP | Position | Allele (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{\mathrm{b}} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $P_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 6 | rs3763311 | 32,376,176 | C/T | 0.27 | 1.13 (1.09-1.17) | $1.17 \times 10^{-10}$ | 1.13 (1.1-1.17) | $5.25 \times 10^{-16}$ | 0.09 | 89\% |
| 6 | rs3763316 | 32,376,746 | C/T | 0.27 | 1.13 (1.09-1.17) | $1.97 \times 10^{-13}$ | 1.13 (1.1-1.16) | $2.95 \times 10^{-18}$ | 0.16 | 96\% |
| 6 | rs9268516 | 32,379,489 | C/T | 0.27 | 1.13 (1.1-1.17) | $4.57 \times 10^{-14}$ | 1.14 (1.1-1.17) | $4.48 \times 10^{-20}$ | 0.10 | 100\% |
| 6 | rs2395163 | 32,387,809 | T/C | 0.20 | 1.12 (1.08-1.17) | $8.02 \times 10^{-9}$ | 1.14 (1.1-1.17) | $1.55 \times 10^{-16}$ | 0.03 | 96\% |
| 6 | rs6926374 | 32,409,305 | A/G | 0.43 | 1.09 (1.06-1.12) | $1.20 \times 10^{-9}$ | 1.09 (1.06-1.12) | $9.26 \times 10^{-12}$ | 0.33 | 96\% |
| 6 | rs2239804 | 32,411,523 | T/C | 0.42 | 1.09 (1.06-1.12) | $1.05 \times 10^{-9}$ | 1.1 (1.07-1.12) | $2.62 \times 10^{-13}$ | 0.21 | 100\% |
| 6 | rs2239803 | 32,411,833 | C/T | 0.43 | 1.09 (1.06-1.12) | $2.26 \times 10^{-9}$ | 1.09 (1.07-1.12) | $4.96 \times 10^{-12}$ | 0.29 | 96\% |
| 6 | rs9268853 | 32,429,643 | T/C | 0.31 | 1.12 (1.08-1.15) | $4.04 \times 10^{-12}$ | 1.12 (1.09-1.15) | $1.72 \times 10^{-16}$ | 0.17 | 98\% |
| 6 | rs9268858 | 32,429,758 | T/C | 0.31 | 1.12 (1.08-1.15) | $3.92 \times 10^{-12}$ | 1.12 (1.09-1.15) | $1.58 \times 10^{-16}$ | 0.17 | 98\% |
| 6 | rs9268877 | 32,431,147 | A/G | 0.57 | 1.09 (1.07-1.12) | $1.94 \times 10^{-11}$ | 1.09 (1.07-1.12) | $1.94 \times 10^{-11}$ | 0.87 | 98\% |
| 6 | rs9268923 | 32,432,835 | C/T | 0.36 | 1.14 (1.11-1.18) | $2.79 \times 10^{-19}$ | 1.14 (1.11-1.18) | $5.37 \times 10^{-22}$ | 0.36 | 93\% |
| 6 | rs2395185 | 32,433,167 | G/T | 0.30 | 1.12 (1.08-1.15) | $3.60 \times 10^{-12}$ | 1.12 (1.09-1.15) | $1.56 \times 10^{-16}$ | 0.17 | 98\% |
| 6 | rs9268969 | 32,434,349 | C/T | 0.36 | 1.14 (1.11-1.18) | $2.65 \times 10^{-19}$ | 1.14 (1.11-1.18) | $5.29 \times 10^{-22}$ | 0.36 | 93\% |
| 6 | rs9268979 | 32,435,044 | T/C | 0.55 | 1.09 (1.06-1.12) | $4.07 \times 10^{-10}$ | 1.09 (1.06-1.12) | $4.07 \times 10^{-10}$ | 0.80 | 95\% |
| 6 | rs9405040 | 32,439,393 | A/C | 0.30 | 1.12 (1.08-1.15) | $4.05 \times 10^{-12}$ | 1.12 (1.09-1.15) | $2.29 \times 10^{-16}$ | 0.17 | 98\% |
| 6 | rs9286790 | 32,439,828 | G/A | 0.30 | 1.12 (1.08-1.15) | $1.55 \times 10^{-12}$ | 1.12 (1.09-1.15) | $3.96 \times 10^{-16}$ | 0.23 | 95\% |
| 6 | rs9269065 | 32,440,172 | T/G | 0.48 | 1.1 (1.07-1.13) | $1.01 \times 10^{-9}$ | 1.1 (1.07-1.13) | $1.01 \times 10{ }^{-9}$ | 0.51 | 84\% |
| 6 | rs9269069 | 32,440,362 | C/T | 0.57 | 1.1 (1.07-1.13) | $1.11 \times 10^{-11}$ | 1.1 (1.07-1.13) | $1.11 \times 10^{-11}$ | 0.85 | 98\% |
| 6 | rs9269070 | 32,440,451 | G/A | 0.58 | 1.09 (1.06-1.12) | $4.08 \times 10^{-9}$ | 1.09 (1.06-1.12) | $4.08 \times 10{ }^{-9}$ | 0.65 | 86\% |
| 6 | rs9269071 | 32,440,467 | T/C | 0.57 | 1.09 (1.06-1.12) | $6.42 \times 10^{-11}$ | 1.09 (1.06-1.12) | $6.42 \times 10^{-11}$ | 0.84 | 95\% |
| 6 | rs9269080 | 32,440,969 | G/A | 0.57 | 1.08 (1.06-1.11) | $1.51 \times 10^{-9}$ | 1.08 (1.06-1.11) | $1.51 \times 10{ }^{-9}$ | 0.69 | 95\% |
| 6 | rs7755212 | 32,441,408 | C/T | 0.57 | 1.09 (1.06-1.12) | $3.22 \times 10^{-10}$ | 1.09 (1.06-1.12) | $3.22 \times 10^{-10}$ | 0.71 | 93\% |
| 6 | rs7739203 | 32,441,555 | A/G | 0.57 | 1.09 (1.06-1.12) | $6.44 \times 10^{-11}$ | 1.09 (1.06-1.12) | $6.44 \times 10^{-11}$ | 0.84 | 95\% |
| 6 | rs7739357 | 32,441,641 | A/G | 0.57 | 1.1 (1.07-1.13) | $4.30 \times 10^{-11}$ | 1.1 (1.07-1.13) | $4.30 \times 10^{-11}$ | 0.82 | 91\% |
| 6 | rs9405112 | 32,445,600 | G/A | 0.30 | 1.12 (1.08-1.15) | $9.54 \times 10^{-12}$ | 1.12 (1.09-1.15) | $1.71 \times 10^{-16}$ | 0.14 | 98\% |
| 6 | rs6916742 | 32,453,191 | T/C | 0.55 | 1.11 (1.08-1.15) | $2.25 \times 10^{-10}$ | 1.12 (1.09-1.15) | $1.23 \times 10^{-14}$ | 0.14 | 93\% |
| 6 | rs2516049 | 32,570,400 | T/C | 0.30 | 1.12 (1.08-1.16) | $5.30 \times 10^{-10}$ | 1.12 (1.09-1.15) | $3.17 \times 10^{-16}$ | 0.03 | 98\% |
| 6 | rs4530903 | 32,581,889 | C/T | 0.12 | 0.87 (0.83-0.91) | $1.85 \times 10^{-9}$ | 0.87 (0.83-0.91) | $1.85 \times 10^{-9}$ | 0.48 | 93\% |
| 6 | rs9272346 | 32,604,372 | G/A | 0.56 | 1.16 (1.13-1.19) | $4.80 \times 10^{-28}$ | 1.16 (1.13-1.19) | $2.35 \times 10^{-28}$ | 0.46 | 96\% |
| 6 | rs9272723 | 32,609,427 | T/C | 0.60 | 1.14 (1.11-1.17) | $1.43 \times 10^{-22}$ | 1.14 (1.11-1.17) | $1.43 \times 10^{-22}$ | 0.68 | 96\% |
| 6 | rs6928482 | 32,626,249 | T/C | 0.45 | 1.11 (1.08-1.15) | $7.28 \times 10^{-11}$ | 1.11 (1.08-1.14) | $3.85 \times 10^{-16}$ | 0.06 | 98\% |
| 6 | rs9273363 | 32,626,272 | C/A | 0.26 | 1.11 (1.07-1.14) | $2.74 \times 10^{-11}$ | 1.11 (1.07-1.14) | $2.74 \times 10^{-11}$ | 0.96 | 96\% |
| 6 | rs6906021 | 32,626,311 | T/C | 0.43 | 1.1 (1.07-1.13) | $1.22 \times 10^{-12}$ | 1.1 (1.07-1.13) | $1.08 \times 10^{-13}$ | 0.39 | 96\% |
| 6 | rs9275141 | 32,651,117 | T/G | 0.47 | 1.11 (1.07-1.14) | $9.66 \times 10^{-10}$ | 1.11 (1.08-1.14) | $5.29 \times 10^{-16}$ | 0.01 | 100\% |
| 6 | rs2858330 | 32,658,715 | T/C | 0.46 | 1.11 (1.07-1.14) | $3.51 \times 10^{-10}$ | 1.11 (1.08-1.14) | $2.35 \times 10^{-16}$ | 0.03 | 100\% |
| 6 | rs17427445 | 32,663,764 | G/A | 0.13 | 0.89 (0.85-0.92) | $6.33 \times 10^{-9}$ | 0.89 (0.85-0.92) | $4.19 \times 10{ }^{-9}$ | 0.43 | 93\% |
| 6 | rs10484561 | 32,665,420 | T/G | 0.13 | 0.89 (0.86-0.93) | $2.81 \times 10^{-9}$ | 0.89 (0.86-0.93) | $2.42 \times 10-9$ | 0.46 | 100\% |
| 6 | rs17427564 | 32,667,067 | T/C | 0.13 | 0.89 (0.85-0.93) | $9.52 \times 10^{-9}$ | 0.89 (0.86-0.93) | $5.26 \times 10^{-9}$ | 0.41 | 95\% |
| 6 | rs10947332 | 32,677,440 | G/A | 0.13 | 0.89 (0.85-0.92) | $2.38 \times 10^{-9}$ | 0.89 (0.85-0.92) | $1.94 \times 10^{-9}$ | 0.45 | 98\% |
| 6 | rs3104405 | 32,682,308 | C/A | 0.65 | 1.11 (1.08-1.14) | $1.61 \times 10^{-12}$ | 1.11 (1.08-1.14) | $1.61 \times 10^{-12}$ | 0.50 | 93\% |
| 6 | rs9275602 | 32,682,812 | C/A | 0.15 | 1.13 (1.09-1.17) | $3.57 \times 10^{-10}$ | 1.13 (1.09-1.17) | $3.57 \times 10^{-10}$ | 0.47 | 91\% |
| 6 | rs241425 | 32,804,909 | A/G | 0.55 | 1.08 (1.05-1.11) | $2.58 \times 10^{-9}$ | 1.08 (1.05-1.11) | $2.58 \times 10^{-9}$ | 0.73 | 98\% |
| 6 | rs2239701 | 32,805,049 | T/C | 0.42 | 1.09 (1.06-1.12) | $1.13 \times 10^{-10}$ | 1.09 (1.06-1.12) | $1.13 \times 10^{-10}$ | 0.95 | 95\% |
| 6 | rs6454802 | 90,814,199 | C/T | 0.37 | 0.93 (0.9-0.95) | $9.05 \times 10^{-9}$ | 0.93 (0.9-0.95) | $9.05 \times 10{ }^{-9}$ | 0.69 | 96\% |
| 6 | rs11753332 | 90,819,153 | G/A | 0.37 | 0.92 (0.9-0.95) | $8.49 \times 10^{-10}$ | 0.92 (0.9-0.95) | $8.49 \times 10^{-10}$ | 0.74 | 100\% |
| 6 | rs12194007 | 90,823,159 | G/T | 0.37 | 0.92 (0.9-0.95) | $2.88 \times 10^{-9}$ | 0.92 (0.9-0.95) | $2.88 \times 10^{-9}$ | 0.66 | 95\% |
| 6 | rs12199079 | 90,852,258 | T/G | 0.34 | 0.92 (0.89-0.94) | $1.20 \times 10^{-10}$ | 0.92 (0.89-0.94) | $1.20 \times 10^{-10}$ | 0.76 | 100\% |
| 6 | rs1010473 | 90,856,878 | G/T | 0.32 | 0.92 (0.89-0.94) | $2.02 \times 10^{-9}$ | 0.92 (0.89-0.94) | $2.02 \times 10{ }^{-9}$ | 0.84 | 93\% |
| 6 | rs1010474 | 90,857,028 | T/C | 0.34 | 0.92 (0.89-0.94) | $1.13 \times 10^{-10}$ | 0.92 (0.89-0.94) | $1.13 \times 10^{-10}$ | 0.71 | 100\% |
| 6 | rs17711850 | 90,864,870 | T/C | 0.34 | 0.92 (0.89-0.94) | $9.03 \times 10^{-11}$ | 0.92 (0.89-0.94) | $9.03 \times 10^{-11}$ | 0.76 | 100\% |
| 6 | rs10455168 | 90,883,525 | T/C | 0.34 | 0.92 (0.89-0.94) | $2.58 \times 10^{-11}$ | 0.92 (0.89-0.94) | $2.58 \times 10^{-11}$ | 0.78 | 100\% |
| 6 | rs10806423 | 90,886,824 | C/T | 0.34 | 0.92 (0.89-0.94) | $2.61 \times 10^{-11}$ | 0.92 (0.89-0.94) | $2.61 \times 10^{-11}$ | 0.78 | 100\% |
| 6 | rs11757155 | 90,941,240 | C/T | 0.33 | 0.91 (0.89-0.94) | $3.07 \times 10^{-12}$ | 0.91 (0.89-0.94) | $3.07 \times 10^{-12}$ | 0.59 | 100\% |
| 6 | rs2021716 | 90,941,289 | C/T | 0.33 | 0.91 (0.89-0.94) | $5.95 \times 10^{-12}$ | 0.91 (0.89-0.94) | $5.95 \times 10^{-12}$ | 0.58 | 100\% |
| 6 | rs17585295 | 90,944,831 | C/T | 0.34 | 0.92 (0.89-0.94) | $1.84 \times 10^{-11}$ | 0.92 (0.89-0.94) | $1.84 \times 10^{-11}$ | 0.67 | 100\% |
| 6 | rs4707609 | 90,946,479 | T/C | 0.34 | 0.92 (0.89-0.94) | $1.85 \times 10^{-11}$ | 0.92 (0.89-0.94) | $1.85 \times 10^{-11}$ | 0.67 | 100\% |
| 6 | rs2875584 | 90,950,628 | C/T | 0.33 | 0.91 (0.89-0.94) | $2.69 \times 10^{-12}$ | 0.91 (0.89-0.94) | $2.69 \times 10^{-12}$ | 0.60 | 100\% |
| 6 | rs17513531 | 90,951,239 | C/T | 0.33 | 0.91 (0.89-0.94) | $2.54 \times 10^{-12}$ | 0.91 (0.89-0.94) | $2.54 \times 10^{-12}$ | 0.61 | 100\% |
| 6 | rs905670 | 90,958,502 | G/A | 0.33 | 0.91 (0.89-0.93) | $1.48 \times 10^{-12}$ | 0.91 (0.89-0.93) | $1.48 \times 10^{-12}$ | 0.68 | 100\% |
| 6 | rs1847472 | 90,973,159 | C/A | 0.33 | 0.91 (0.89-0.94) | $6.95 \times 10^{-12}$ | 0.91 (0.89-0.94) | $6.95 \times 10^{-12}$ | 0.68 | 98\% |


| Chrom | SNP | Position | Allele (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 6 | rs943689 | 90,984,035 | C/T | 0.33 | 0.91 (0.89-0.93) | $1.23 \times 10^{-12}$ | 0.91 (0.89-0.93) | $1.23 \times 10^{-12}$ | 0.75 | 100\% |
| 6 | rs2325291 | 90,986,686 | G/A | 0.33 | 0.91 (0.89-0.93) | $8.58 \times 10^{-13}$ | 0.91 (0.89-0.93) | $8.58 \times 10^{-13}$ | 0.78 | 100\% |
| 6 | rs2325292 | 90,986,749 | T/C | 0.33 | 0.91 (0.89-0.93) | $9.00 \times 10^{-13}$ | 0.91 (0.89-0.93) | $9.00 \times 10^{-13}$ | 0.78 | 100\% |
| 6 | rs2174281 | 90,987,872 | T/C | 0.45 | 0.92 (0.9-0.94) | $7.78 \times 10^{-11}$ | 0.92 (0.9-0.94) | $7.78 \times 10^{-11}$ | 0.51 | 100\% |
| 6 | rs4142967 | 90,996,349 | C/T | 0.45 | 0.92 (0.9-0.95) | $1.54 \times 10^{-10}$ | 0.92 (0.9-0.95) | $1.54 \times 10^{-10}$ | 0.49 | 100\% |
| 6 | rs12212193 | 90,996,769 | A/G | 0.45 | 0.92 (0.9-0.94) | $8.42 \times 10^{-11}$ | 0.92 (0.9-0.94) | $8.42 \times 10^{-11}$ | 0.50 | 100\% |
| 6 | rs1504215 | 91,006,227 | G/A | 0.33 | 0.91 (0.89-0.93) | $1.77 \times 10^{-12}$ | 0.91 (0.89-0.93) | $1.77 \times 10^{-12}$ | 0.81 | 100\% |
| 6 | rs6925032 | 91,008,027 | C/A | 0.33 | 0.91 (0.89-0.93) | $2.04 \times 10^{-12}$ | 0.91 (0.89-0.93) | $2.04 \times 10^{-12}$ | 0.81 | 100\% |
| 6 | rs1321859 | 91,011,673 | C/T | 0.33 | 0.91 (0.89-0.93) | $2.58 \times 10^{-12}$ | 0.91 (0.89-0.93) | $2.58 \times 10^{-12}$ | 0.81 | 100\% |
| 9 | rs10975413 | 6,049,843 | A/G | 0.19 | 0.87 (0.83-0.91) | $7.69 \times 10^{-9}$ | 0.9 (0.87-0.93) | $1.26 \times 10^{-10}$ | 0.01 | 96\% |
| 9 | rs380568 | 6,055,531 | T/C | 0.19 | 0.88 (0.84-0.92) | $9.43 \times 10^{-9}$ | 0.9 (0.87-0.93) | $1.85 \times 10^{-10}$ | 0.00 | 100\% |
| 9 | rs343490 | 6,064,575 | A/G | 0.19 | 0.88 (0.84-0.92) | $6.75 \times 10^{-9}$ | 0.9 (0.87-0.93) | $1.10 \times 10^{-10}$ | 0.00 | 100\% |
| 9 | rs343476 | 6,072,597 | T/C | 0.82 | 1.16 (1.1-1.22) | $9.15 \times 10^{-9}$ | 1.12 (1.08-1.16) | $1.01 \times 10^{-10}$ | 0.00 | 98\% |
| 9 | rs378952 | 6,078,146 | C/T | 0.81 | 1.14 (1.09-1.19) | $5.55 \times 10^{-9}$ | 1.11 (1.08-1.15) | $4.91 \times 10^{-11}$ | 0.01 | 100\% |
| 9 | rs371454 | 6,078,614 | C/T | 0.81 | 1.14 (1.09-1.19) | $6.45 \times 10^{-9}$ | 1.11 (1.08-1.15) | $5.63 \times 10^{-11}$ | 0.01 | 100\% |
| 9 | rs413382 | 6,142,948 | C/A | 0.81 | 1.14 (1.09-1.19) | $3.47 \times 10^{-9}$ | 1.12 (1.08-1.16) | $9.31 \times 10^{-12}$ | 0.01 | 98\% |
| 9 | rs369756 | 6,146,441 | G/T | 0.23 | 1.12 (1.09-1.16) | $5.36 \times 10^{-12}$ | 1.12 (1.09-1.16) | $2.75 \times 10^{-13}$ | 0.34 | 95\% |
| 9 | rs450108 | 6,153,485 | T/C | 0.42 | 1.1 (1.07-1.14) | $6.46 \times 10^{-9}$ | 1.1 (1.07-1.13) | $2.29 \times 10^{-11}$ | 0.14 | 89\% |
| 9 | rs1116795 | 6,155,226 | G/T | 0.41 | 1.1 (1.07-1.13) | $9.99 \times 10^{-10}$ | 1.09 (1.07-1.12) | $4.06 \times 10^{-12}$ | 0.16 | 98\% |
| 9 | rs2225537 | 6,160,578 | C/T | 0.41 | 1.1 (1.07-1.13) | $8.94 \times 10^{-10}$ | 1.09 (1.07-1.12) | $2.69 \times 10^{-12}$ | 0.14 | 98\% |
| 9 | rs10124250 | 6,161,686 | C/T | 0.41 | 1.1 (1.07-1.13) | $9.61 \times 10^{-10}$ | 1.09 (1.07-1.12) | $2.40 \times 10^{-12}$ | 0.14 | 98\% |
| 9 | rs10119713 | 6,163,823 | G/A | 0.41 | 1.1 (1.07-1.13) | $7.97 \times 10^{-10}$ | 1.09 (1.07-1.12) | $2.42 \times 10^{-12}$ | 0.14 | 98\% |
| 9 | rs7032572 | 6,172,380 | A/G | 0.16 | 1.18 (1.13-1.23) | $1.24 \times 10^{-14}$ | 1.18 (1.14-1.22) | $3.85 \times 10^{-23}$ | 0.03 | 100\% |
| 9 | rs1412426 | 6,188,652 | A/C | 0.68 | 0.9 (0.86-0.93) | $3.20 \times 10^{-9}$ | 0.9 (0.87-0.92) | $9.42 \times 10^{-17}$ | 0.00 | 100\% |
| 9 | rs1412425 | 6,188,740 | A/C | 0.68 | 0.9 (0.86-0.93) | $1.51 \times 10^{-9}$ | 0.89 (0.87-0.92) | $3.48 \times 10^{-17}$ | 0.00 | 100\% |
| 9 | rs1342326 | 6,190,076 | A/C | 0.16 | 1.18 (1.13-1.23) | $4.68 \times 10^{-15}$ | 1.18 (1.15-1.22) | $2.11 \times 10^{-24}$ | 0.03 | 100\% |
| 9 | rs2095044 | 6,192,796 | T/C | 0.73 | 0.86 (0.83-0.89) | $4.09 \times 10^{-16}$ | 0.86 (0.84-0.89) | $3.12 \times 10^{-26}$ | 0.01 | 100\% |
| 9 | rs2381416 | 6,193,455 | C/A | 0.73 | 0.85 (0.82-0.89) | $1.23 \times 10^{-16}$ | 0.86 (0.84-0.88) | $4.54 \times 10^{-27}$ | 0.01 | 100\% |
| 9 | rs10815370 | 6,194,831 | C/A | 0.66 | 0.89 (0.86-0.93) | $3.06 \times 10^{-9}$ | 0.89 (0.87-0.92) | $9.99 \times 10^{-17}$ | 0.00 | 98\% |
| 9 | rs1888909 | 6,197,392 | T/C | 0.73 | 0.86 (0.82-0.89) | $3.26 \times 10^{-16}$ | 0.86 (0.84-0.88) | $9.52 \times 10^{-27}$ | 0.01 | 100\% |
| 9 | rs992969 | 6,209,697 | A/G | 0.75 | 0.85 (0.82-0.88) | $1.08 \times 10^{-17}$ | 0.85 (0.83-0.88) | $4.27 \times 10^{-29}$ | 0.01 | 100\% |
| 9 | rs3939286 | 6,210,099 | T/C | 0.75 | 0.85 (0.82-0.88) | $2.24 \times 10^{-17}$ | 0.86 (0.83-0.88) | $2.29 \times 10^{-28}$ | 0.01 | 100\% |
| 9 | rs928413 | 6,213,387 | G/A | 0.74 | 0.85 (0.82-0.88) | $5.44 \times 10^{-16}$ | 0.86 (0.83-0.88) | $1.79 \times 10^{-27}$ | 0.00 | 98\% |
| 9 | rs7848215 | 6,213,468 | C/T | 0.26 | 1.18 (1.13-1.23) | $3.33 \times 10^{-17}$ | 1.17 (1.14-1.2) | $3.08 \times 10^{-28}$ | 0.01 | 98\% |
| 9 | rs2066362 | 6,219,176 | G/T | 0.16 | 1.18 (1.13-1.24) | $1.07 \times 10^{-14}$ | 1.18 (1.14-1.22) | $1.54 \times 10^{-22}$ | 0.05 | 98\% |
| 9 | rs12551256 | 6,231,239 | A/G | 0.47 | 0.92 (0.89-0.94) | $1.69 \times 10^{-9}$ | 0.92 (0.89-0.94) | $2.15 \times 10^{-11}$ | 0.26 | 96\% |
| 9 | rs17582919 | 6,233,376 | T/C | 0.22 | 1.13 (1.09-1.17) | $1.22 \times 10^{-10}$ | 1.13 (1.1-1.16) | $6.73 \times 10^{-16}$ | 0.04 | 100\% |
| 9 | rs17498196 | 6,237,547 | A/C | 0.22 | 1.13 (1.09-1.17) | $8.84 \times 10^{-11}$ | 1.13 (1.09-1.16) | $1.45 \times 10^{-15}$ | 0.06 | 100\% |
| 9 | rs10815393 | 6,240,324 | T/C | 0.22 | 1.12 (1.08-1.17) | $4.42 \times 10^{-9}$ | 1.12 (1.09-1.16) | $7.06 \times 10^{-14}$ | 0.04 | 95\% |
| 11 | rs2155219 | 76,299,194 | G/T | 0.46 | 1.11 (1.08-1.15) | $2.43 \times 10^{-10}$ | 1.11 (1.08-1.14) | $2.90 \times 10^{-15}$ | 0.04 | 95\% |
| 11 | rs7927894 | 76,301,316 | C/T | 0.37 | 1.1 (1.07-1.13) | $3.52 \times 10^{-11}$ | 1.1 (1.07-1.13) | $3.23 \times 10^{-12}$ | 0.38 | 95\% |
| 11 | rs7927997 | 76,301,375 | C/T | 0.37 | 1.1 (1.07-1.13) | $3.91 \times 10^{-11}$ | 1.1 (1.07-1.13) | $8.40 \times 10^{-12}$ | 0.41 | 91\% |
| 15 | rs10519067 | 61,068,347 | G/A | 0.14 | 0.89 (0.85-0.92) | $8.28 \times 10^{-9}$ | 0.89 (0.85-0.92) | $1.49 \times 10^{-10}$ | 0.30 | 100\% |
| 15 | rs11071558 | 61,069,421 | A/G | 0.14 | 0.89 (0.85-0.92) | $1.85 \times 10^{-10}$ | 0.89 (0.85-0.92) | $8.32 \times 10^{-11}$ | 0.44 | 100\% |
| 15 | rs11071559 | 61,069,988 | C/T | 0.14 | 0.89 (0.86-0.92) | $1.93 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.93 \times 10^{-10}$ | 0.52 | 100\% |
| 15 | rs1866316 | 67,441,997 | T/C | 0.29 | 1.12 (1.08-1.15) | $3.03 \times 10^{-14}$ | 1.12 (1.08-1.15) | $3.03 \times 10^{-14}$ | 0.80 | 93\% |
| 15 | rs17293632 | 67,442,596 | C/T | 0.26 | 1.12 (1.09-1.16) | $5.10 \times 10^{-14}$ | 1.13 (1.09-1.16) | $8.81 \times 10^{-16}$ | 0.36 | 98\% |
| 15 | rs10152544 | 67,444,747 | C/T | 0.50 | 0.91 (0.89-0.94) | $1.73 \times 10^{-12}$ | 0.91 (0.89-0.94) | $1.73 \times 10^{-12}$ | 0.97 | 96\% |
| 15 | rs744910 | 67,446,785 | G/A | 0.50 | 0.92 (0.89-0.94) | $9.76 \times 10^{-13}$ | 0.92 (0.89-0.94) | $9.76 \times 10^{-13}$ | 0.98 | 100\% |
| 15 | rs11634793 | 67,447,452 | C/T | 0.50 | 0.92 (0.89-0.94) | $1.56 \times 10^{-12}$ | 0.92 (0.89-0.94) | $1.56 \times 10^{-12}$ | 0.98 | 100\% |
| 15 | rs8032739 | 67,448,899 | A/G | 0.30 | 1.11 (1.08-1.14) | $5.58 \times 10^{-14}$ | 1.11 (1.08-1.14) | $5.58 \times 10^{-14}$ | 0.77 | 96\% |
| 15 | rs2033784 | 67,449,660 | A/G | 0.30 | 1.11 (1.08-1.14) | $2.47 \times 10^{-14}$ | 1.11 (1.08-1.14) | $2.47 \times 10^{-14}$ | 0.75 | 100\% |
| 15 | rs17228058 | 67,450,305 | A/G | 0.26 | 1.12 (1.09-1.16) | $1.32 \times 10^{-12}$ | 1.12 (1.09-1.16) | $2.85 \times 10^{-15}$ | 0.27 | 96\% |
| 15 | rs7173698 | 67,450,893 | A/G | 0.30 | 1.11 (1.08-1.14) | $5.83 \times 10^{-14}$ | 1.11 (1.08-1.14) | $5.83 \times 10^{-14}$ | 0.76 | 96\% |
| 15 | rs4562997 | 67,458,152 | G/A | 0.29 | 1.11 (1.08-1.14) | $2.04 \times 10^{-13}$ | 1.11 (1.08-1.14) | $2.04 \times 10^{-13}$ | 0.71 | 100\% |
| 15 | rs16950687 | 67,464,013 | A/G | 0.28 | 1.1 (1.07-1.13) | $5.28 \times 10^{-12}$ | 1.1 (1.07-1.13) | $5.28 \times 10^{-12}$ | 0.79 | 100\% |
| 15 | rs12708492 | 67,467,541 | T/C | 0.51 | 0.92 (0.9-0.94) | $1.88 \times 10^{-11}$ | 0.92 (0.9-0.94) | $1.88 \times 10^{-11}$ | 0.85 | 100\% |
| 15 | rs17294280 | 67,468,285 | A/G | 0.27 | 1.12 (1.08-1.17) | $4.07 \times 10^{-10}$ | 1.13 (1.09-1.16) | $3.76 \times 10^{-14}$ | 0.15 | 96\% |
| 16 | rs3743976 | 11,038,824 | C/T | 0.22 | 0.91 (0.88-0.94) | $4.57 \times 10^{-9}$ | 0.9 (0.88-0.93) | $1.78 \times 10^{-10}$ | 0.36 | 100\% |
| 16 | rs8062923 | 11,160,966 | A/C | 0.76 | 1.1 (1.07-1.14) | $1.45 \times 10^{-9}$ | 1.11 (1.08-1.14) | $7.84 \times 10^{-12}$ | 0.28 | 100\% |


| Chrom | SNP | Position | Allele <br> (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P h e t}^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 16 | rs4781035 | 11,161,178 | A/G | 0.79 | 1.1 (1.07-1.14) | $8.98 \times 10^{-10}$ | 1.1 (1.07-1.14) | $2.81 \times 10^{-10}$ | 0.43 | 100\% |
| 16 | rs7192790 | 11,167,914 | C/A | 0.20 | 0.91 (0.88-0.94) | $3.65 \times 10^{-9}$ | 0.91 (0.88-0.94) | $3.65 \times 10{ }^{-9}$ | 0.52 | 100\% |
| 16 | rs12444495 | 11,170,455 | T/C | 0.79 | 1.1 (1.07-1.14) | $1.12 \times 10^{-9}$ | 1.1 (1.07-1.14) | $6.93 \times 10^{-10}$ | 0.46 | 95\% |
| 16 | rs7184491 | 11,170,761 | C/T | 0.78 | 1.1 (1.07-1.14) | $6.74 \times 10^{-9}$ | 1.1 (1.07-1.14) | $2.03 \times 10^{-9}$ | 0.43 | 89\% |
| 16 | rs17806299 | 11,199,980 | G/A | 0.20 | 0.9 (0.88-0.93) | $2.09 \times 10^{-10}$ | 0.9 (0.88-0.93) | $2.09 \times 10^{-10}$ | 0.51 | 100\% |
| 16 | rs12935657 | 11,219,041 | G/A | 0.25 | 0.91 (0.88-0.94) | $3.71 \times 10^{-9}$ | 0.9 (0.88-0.93) | $2.06 \times 10^{-12}$ | 0.22 | 98\% |
| 16 | rs7203459 | 11,230,703 | T/C | 0.26 | 0.91 (0.88-0.94) | $8.79 \times 10^{-9}$ | 0.9 (0.88-0.93) | $5.59 \times 10^{-12}$ | 0.22 | 95\% |
| 17 | rs12950186 | 37,393,395 | A/C | 0.85 | 0.9 (0.87-0.93) | $2.03 \times 10^{-9}$ | 0.9 (0.87-0.93) | $2.03 \times 10^{-9}$ | 0.84 | 98\% |
| 17 | rs2879258 | 37,399,379 | G/T | 0.85 | 0.9 (0.87-0.93) | $1.93 \times 10^{-9}$ | 0.9 (0.87-0.93) | $1.93 \times 10^{-9}$ | 0.88 | 100\% |
| 17 | rs11078895 | 37,401,051 | A/G | 0.85 | 0.9 (0.87-0.93) | $1.96 \times 10^{-9}$ | 0.9 (0.87-0.93) | $1.96 \times 10^{-9}$ | 0.89 | 98\% |
| 17 | rs11655972 | 37,407,072 | C/T | 0.85 | 0.9 (0.87-0.93) | $8.02 \times 10^{-10}$ | 0.9 (0.87-0.93) | $8.02 \times 10^{-10}$ | 0.90 | 100\% |
| 17 | rs3744349 | 37,414,842 | C/T | 0.85 | 0.9 (0.87-0.93) | $7.20 \times 10^{-10}$ | 0.9 (0.87-0.93) | $7.20 \times 10^{-10}$ | 0.91 | 100\% |
| 17 | rs8073907 | 37,424,149 | C/T | 0.85 | 0.9 (0.87-0.93) | $1.60 \times 10^{-9}$ | 0.9 (0.87-0.93) | $1.60 \times 10^{-9}$ | 0.93 | 95\% |
| 17 | rs667239 | 37,442,241 | A/G | 0.15 | 1.11 (1.07-1.15) | $6.49 \times 10^{-10}$ | 1.11 (1.07-1.15) | $6.49 \times 10^{-10}$ | 0.93 | 100\% |
| 17 | rs590051 | 37,446,571 | T/C | 0.15 | 1.11 (1.07-1.15) | $6.53 \times 10^{-10}$ | 1.11 (1.07-1.15) | $6.53 \times 10^{-10}$ | 0.93 | 100\% |
| 17 | rs2302073 | 37,457,342 | A/G | 0.15 | 1.11 (1.08-1.15) | $5.94 \times 10^{-10}$ | 1.11 (1.08-1.15) | $5.94 \times 10^{-10}$ | 0.93 | 100\% |
| 17 | rs584377 | 37,460,128 | G/A | 0.15 | 1.11 (1.07-1.15) | $6.04 \times 10^{-10}$ | 1.11 (1.07-1.15) | $6.04 \times 10^{-10}$ | 0.93 | 100\% |
| 17 | rs649180 | 37,464,959 | A/C | 0.15 | 1.11 (1.07-1.15) | $6.49 \times 10^{-10}$ | 1.11 (1.07-1.15) | $6.49 \times 10^{-10}$ | 0.93 | 100\% |
| 17 | rs2338799 | 37,513,941 | G/A | 0.15 | 1.11 (1.07-1.15) | $8.15 \times 10^{-10}$ | 1.11 (1.07-1.15) | $8.15 \times 10^{-10}$ | 0.94 | 100\% |
| 17 | rs6503504 | 37,514,412 | A/G | 0.15 | 1.11 (1.07-1.15) | $1.29 \times 10^{-9}$ | 1.11 (1.07-1.15) | $1.29 \times 10^{-9}$ | 0.92 | 96\% |
| 17 | rs9908131 | 37,520,449 | T/C | 0.15 | 1.11 (1.07-1.15) | $8.73 \times 10^{-10}$ | 1.11 (1.07-1.15) | $8.73 \times 10^{-10}$ | 0.94 | 100\% |
| 17 | rs7208487 | 37,543,449 | T/G | 0.15 | 1.11 (1.07-1.15) | $8.79 \times 10^{-10}$ | 1.11 (1.07-1.15) | $8.79 \times 10^{-10}$ | 0.94 | 100\% |
| 17 | rs4795358 | 37,573,065 | A/C | 0.85 | 0.9 (0.87-0.94) | $6.62 \times 10^{-9}$ | 0.9 (0.87-0.94) | $6.62 \times 10^{-9}$ | 0.96 | 96\% |
| 17 | rs9646419 | 37,597,185 | A/G | 0.85 | 0.9 (0.87-0.94) | $6.48 \times 10^{-9}$ | 0.9 (0.87-0.94) | $6.48 \times 10^{-9}$ | 0.96 | 96\% |
| 17 | rs12938099 | 37,612,910 | T/G | 0.85 | 0.9 (0.87-0.94) | $5.14 \times 10^{-9}$ | 0.9 (0.87-0.94) | $5.14 \times 10^{-9}$ | 0.96 | 96\% |
| 17 | rs12937013 | 37,665,571 | G/A | 0.85 | 0.9 (0.87-0.93) | $2.35 \times 10^{-10}$ | 0.9 (0.87-0.93) | $2.35 \times 10^{-10}$ | 0.95 | 100\% |
| 17 | rs8069074 | 37,685,401 | A/G | 0.85 | 0.9 (0.87-0.93) | $1.54 \times 10^{-9}$ | 0.9 (0.87-0.93) | $1.54 \times 10^{-9}$ | 0.93 | 98\% |
| 17 | rs11657058 | 37,699,378 | T/G | 0.85 | 0.9 (0.87-0.93) | $3.18 \times 10^{-10}$ | 0.9 (0.87-0.93) | $3.18 \times 10^{-10}$ | 0.95 | 100\% |
| 17 | rs11657153 | 37,699,729 | A/G | 0.85 | 0.9 (0.87-0.93) | $5.95 \times 10^{-10}$ | 0.9 (0.87-0.93) | $5.95 \times 10^{-10}$ | 0.94 | 95\% |
| 17 | rs12947506 | 37,707,592 | T/C | 0.85 | 0.9 (0.87-0.93) | $3.83 \times 10^{-10}$ | 0.9 (0.87-0.93) | $3.83 \times 10^{-10}$ | 0.95 | 100\% |
| 17 | rs7503377 | 37,708,841 | T/C | 0.85 | 0.9 (0.87-0.93) | $3.71 \times 10^{-10}$ | 0.9 (0.87-0.93) | $3.71 \times 10^{-10}$ | 0.95 | 100\% |
| 17 | rs6503521 | 37,715,551 | G/T | 0.85 | 0.89 (0.86-0.93) | $1.54 \times 10^{-10}$ | 0.89 (0.86-0.93) | $1.54 \times 10^{-10}$ | 0.95 | 98\% |
| 17 | rs903507 | 37,726,423 | C/T | 0.15 | 1.12 (1.08-1.16) | $3.90 \times 10^{-11}$ | 1.12 (1.08-1.16) | $3.90 \times 10^{-11}$ | 0.94 | 98\% |
| 17 | rs8182252 | 37,727,950 | T/C | 0.17 | 1.11 (1.07-1.15) | $4.28 \times 10^{-9}$ | 1.11 (1.07-1.15) | $4.28 \times 10^{-9}$ | 0.79 | 93\% |
| 17 | rs1874226 | 37,729,031 | C/T | 0.15 | 1.12 (1.09-1.16) | $2.64 \times 10^{-11}$ | 1.12 (1.09-1.16) | $2.64 \times 10^{-11}$ | 0.94 | 98\% |
| 17 | rs4795385 | 37,733,148 | A/G | 0.17 | 1.13 (1.09-1.17) | $1.16 \times 10^{-12}$ | 1.13 (1.09-1.17) | $1.16 \times 10^{-12}$ | 0.92 | 96\% |
| 17 | rs1877030 | 37,740,161 | T/C | 0.84 | 0.89 (0.86-0.92) | $1.80 \times 10^{-12}$ | 0.89 (0.86-0.92) | $1.80 \times 10^{-12}$ | 0.97 | 98\% |
| 17 | rs12453198 | 37,741,879 | T/C | 0.84 | 0.88 (0.85-0.91) | $3.48 \times 10^{-14}$ | 0.88 (0.85-0.91) | $3.48 \times 10^{-14}$ | 0.97 | 98\% |
| 17 | rs11654954 | 37,745,979 | A/G | 0.83 | 0.88 (0.85-0.91) | $1.57 \times 10^{-14}$ | 0.88 (0.85-0.91) | $1.57 \times 10^{-14}$ | 0.95 | 98\% |
| 17 | rs12453682 | 37,770,005 | C/T | 0.70 | 0.9 (0.88-0.93) | $7.00 \times 10^{-12}$ | 0.9 (0.88-0.93) | $9.56 \times 10^{-14}$ | 0.29 | 98\% |
| 17 | rs1874228 | 37,775,274 | A/G | 0.73 | 0.91 (0.88-0.93) | $5.81 \times 10^{-12}$ | 0.91 (0.88-0.93) | $5.81 \times 10^{-12}$ | 0.56 | 96\% |
| 17 | rs879606 | 37,781,849 | A/G | 0.83 | 0.88 (0.85-0.91) | $1.61 \times 10^{-13}$ | 0.88 (0.85-0.91) | $1.61 \times 10^{-13}$ | 0.97 | 95\% |
| 17 | rs2271309 | 37,784,990 | G/A | 0.80 | 0.89 (0.86-0.92) | $4.27 \times 10^{-14}$ | 0.89 (0.86-0.92) | $4.27 \times 10^{-14}$ | 0.92 | 98\% |
| 17 | rs907094 | 37,790,371 | G/A | 0.80 | 0.89 (0.87-0.92) | $1.28 \times 10^{-12}$ | 0.89 (0.87-0.92) | $1.28 \times 10^{-12}$ | 0.89 | 98\% |
| 17 | rs3764352 | 37,790,939 | C/T | 0.80 | 0.9 (0.87-0.93) | $4.64 \times 10^{-10}$ | 0.9 (0.87-0.93) | $4.64 \times 10^{-10}$ | 0.88 | 88\% |
| 17 | rs9972882 | 37,807,698 | A/C | 0.75 | 0.88 (0.85-0.9) | $1.85 \times 10^{-20}$ | 0.88 (0.85-0.9) | $1.85 \times 10^{-20}$ | 0.66 | 98\% |
| 17 | rs1877031 | 37,814,080 | G/A | 0.67 | 0.89 (0.87-0.91) | $1.32 \times 10^{-18}$ | 0.89 (0.87-0.91) | $1.32 \times 10^{-18}$ | 0.58 | 98\% |
| 17 | rs2271308 | 37,817,482 | T/C | 0.73 | 0.89 (0.86-0.91) | $1.93 \times 10^{-17}$ | 0.89 (0.86-0.91) | $1.93 \times 10^{-17}$ | 0.69 | 98\% |
| 17 | rs931992 | 37,821,435 | G/T | 0.67 | 0.89 (0.87-0.91) | $2.55 \times 10^{-18}$ | 0.89 (0.87-0.91) | $2.55 \times 10^{-18}$ | 0.56 | 98\% |
| 17 | rs1053651 | 37,822,311 | A/C | 0.73 | 0.89 (0.86-0.92) | $1.22 \times 10^{-15}$ | 0.89 (0.86-0.92) | $1.22 \times 10^{-15}$ | 0.59 | 93\% |
| 17 | rs876493 | 37,824,545 | G/A | 0.60 | 0.88 (0.85-0.91) | $1.17 \times 10^{-11}$ | 0.88 (0.85-0.91) | $1.92 \times 10^{-16}$ | 0.13 | 82\% |
| 17 | rs14050 | 37,828,072 | C/T | 0.70 | 0.86 (0.84-0.89) | $8.89 \times 10^{-23}$ | 0.86 (0.84-0.89) | $8.89 \times 10^{-23}$ | 0.49 | 88\% |
| 17 | rs2952151 | 37,828,496 | T/C | 0.70 | 0.86 (0.84-0.89) | $2.32 \times 10^{-27}$ | 0.86 (0.84-0.89) | $2.32 \times 10^{-27}$ | 0.66 | 98\% |
| 17 | rs2941503 | 37,828,745 | A/G | 0.70 | 0.87 (0.84-0.89) | $6.42 \times 10^{-23}$ | 0.87 (0.84-0.89) | $6.42 \times 10^{-23}$ | 0.53 | 89\% |
| 17 | rs907087 | 37,828,787 | G/A | 0.70 | 0.86 (0.84-0.89) | $6.67 \times 10^{-23}$ | 0.86 (0.84-0.89) | $6.67 \times 10^{-23}$ | 0.49 | 88\% |
| 17 | rs903502 | 37,829,604 | C/T | 0.68 | 0.87 (0.85-0.9) | $1.43 \times 10^{-24}$ | 0.87 (0.85-0.9) | $1.43 \times 10^{-24}$ | 0.62 | 100\% |
| 17 | rs2941504 | 37,830,900 | A/G | 0.70 | 0.86 (0.84-0.89) | $1.28 \times 10^{-27}$ | 0.86 (0.84-0.89) | $1.28 \times 10^{-27}$ | 0.65 | 98\% |
| 17 | rs1565922 | 37,831,035 | A/G | 0.70 | 0.86 (0.84-0.89) | $3.85 \times 10^{-27}$ | 0.86 (0.84-0.89) | $3.85 \times 10^{-27}$ | 0.65 | 98\% |
| 17 | rs2934952 | 37,832,366 | G/A | 0.71 | 0.87 (0.84-0.89) | $4.33 \times 10^{-24}$ | 0.87 (0.84-0.89) | $4.33 \times 10^{-24}$ | 0.49 | 89\% |
| 17 | rs2941505 | 37,832,704 | A/G | 0.68 | 0.87 (0.85-0.9) | $2.83 \times 10^{-24}$ | 0.87 (0.85-0.9) | $2.83 \times 10^{-24}$ | 0.51 | 100\% |


| Chrom | SNP | Position | Allele (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 17 | rs2941506 | 37,833,035 | A/G | 0.68 | 0.87 (0.85-0.9) | $2.21 \times 10^{-22}$ | 0.87 (0.85-0.9) | $2.17 \times 10^{-23}$ | 0.43 | 96\% |
| 17 | rs907089 | 37,833,600 | G/A | 0.68 | 0.88 (0.85-0.9) | $1.42 \times 10^{-22}$ | 0.88 (0.85-0.9) | $1.42 \times 10^{-22}$ | 0.51 | 98\% |
| 17 | rs2313171 | 37,833,842 | T/C | 0.68 | 0.87 (0.85-0.9) | $3.82 \times 10^{-22}$ | 0.87 (0.85-0.9) | $2.50 \times 10^{-23}$ | 0.42 | 96\% |
| 17 | rs12150298 | 37,834,541 | T/C | 0.68 | 0.88 (0.86-0.9) | $9.04 \times 10^{-23}$ | 0.88 (0.86-0.9) | $4.46 \times 10^{-23}$ | 0.46 | 100\% |
| 17 | rs8078228 | 37,834,998 | C/T | 0.68 | 0.87 (0.85-0.9) | $5.04 \times 10^{-22}$ | 0.87 (0.85-0.9) | $2.05 \times 10^{-23}$ | 0.41 | 96\% |
| 17 | rs11078919 | 37,835,755 | T/C | 0.68 | 0.88 (0.85-0.9) | $9.43 \times 10^{-20}$ | 0.88 (0.86-0.9) | $2.59 \times 10^{-21}$ | 0.39 | 95\% |
| 17 | rs1476278 | 37,836,243 | C/T | 0.68 | 0.88 (0.85-0.9) | $7.34 \times 10^{-23}$ | 0.88 (0.85-0.9) | $2.15 \times 10^{-23}$ | 0.45 | 100\% |
| 17 | rs9303274 | 37,836,353 | T/C | 0.68 | 0.88 (0.85-0.9) | $7.40 \times 10^{-21}$ | 0.88 (0.85-0.9) | $1.44 \times 10^{-22}$ | 0.39 | 96\% |
| 17 | rs2517957 | 37,838,716 | G/A | 0.68 | 0.87 (0.85-0.9) | $4.64 \times 10^{-24}$ | 0.87 (0.85-0.9) | $3.13 \times 10^{-24}$ | 0.47 | 100\% |
| 17 | rs2517958 | 37,838,751 | G/A | 0.68 | 0.87 (0.85-0.9) | $5.66 \times 10^{-24}$ | 0.87 (0.85-0.9) | $3.07 \times 10^{-24}$ | 0.46 | 100\% |
| 17 | rs903501 | 37,839,493 | T/C | 0.70 | 0.87 (0.84-0.9) | $1.55 \times 10^{-19}$ | 0.87 (0.84-0.9) | $2.68 \times 10^{-21}$ | 0.40 | 88\% |
| 17 | rs2517954 | 37,843,550 | T/C | 0.69 | 0.86 (0.84-0.88) | $4.84 \times 10^{-28}$ | 0.86 (0.84-0.88) | $4.84 \times 10^{-28}$ | 0.63 | 98\% |
| 17 | rs2517955 | 37,843,681 | C/T | 0.67 | 0.87 (0.85-0.9) | $2.22 \times 10^{-23}$ | 0.87 (0.85-0.9) | $2.22 \times 10^{-23}$ | 0.53 | 98\% |
| 17 | rs2517956 | 37,843,859 | G/A | 0.69 | 0.86 (0.84-0.89) | $1.05 \times 10^{-26}$ | 0.86 (0.84-0.89) | $1.05 \times 10^{-26}$ | 0.63 | 98\% |
| 17 | rs1565923 | 37,858,678 | A/G | 0.70 | 0.86 (0.84-0.89) | $2.09 \times 10^{-25}$ | 0.86 (0.84-0.89) | $2.09 \times 10^{-25}$ | 0.50 | 93\% |
| 17 | rs2952155 | 37,861,718 | T/C | 0.76 | 0.86 (0.83-0.88) | $2.80 \times 10^{-24}$ | 0.86 (0.83-0.88) | $2.80 \times 10^{-24}$ | 0.76 | 98\% |
| 17 | rs1810132 | 37,866,005 | C/T | 0.70 | 0.86 (0.84-0.88) | $7.92 \times 10^{-29}$ | 0.86 (0.84-0.88) | $7.92 \times 10^{-29}$ | 0.54 | 98\% |
| 17 | rs2952156 | 37,876,835 | A/G | 0.70 | 0.86 (0.84-0.88) | $7.55 \times 10^{-29}$ | 0.86 (0.84-0.88) | $7.55 \times 10^{-29}$ | 0.55 | 98\% |
| 17 | rs907091 | 37,921,742 | C/T | 0.48 | 1.18 (1.14-1.22) | $2.00 \times 10^{-20}$ | 1.19 (1.15-1.22) | $1.12 \times 10^{-36}$ | 0.01 | 95\% |
| 17 | rs907092 | 37,922,259 | G/A | 0.48 | 0.86 (0.83-0.89) | $1.55 \times 10^{-18}$ | 0.85 (0.83-0.88) | $2.16 \times 10^{-32}$ | 0.01 | 96\% |
| 17 | rs10445308 | 37,938,047 | C/T | 0.48 | 0.85 (0.83-0.88) | $6.58 \times 10^{-19}$ | 0.85 (0.83-0.87) | $3.99 \times 10^{-34}$ | 0.01 | 98\% |
| 17 | rs9909593 | 37,970,149 | A/G | 0.48 | 0.86 (0.83-0.89) | $1.14 \times 10^{-18}$ | 0.85 (0.83-0.88) | $4.16 \times 10^{-34}$ | 0.01 | 98\% |
| 17 | rs9303277 | 37,976,469 | C/T | 0.52 | 0.84 (0.82-0.87) | $5.43 \times 10^{-22}$ | 0.84 (0.82-0.86) | $1.18 \times 10^{-40}$ | 0.01 | 98\% |
| 17 | rs3816470 | 37,985,801 | A/G | 0.53 | 0.85 (0.82-0.88) | $1.23 \times 10^{-21}$ | 0.84 (0.82-0.86) | $2.79 \times 10^{-38}$ | 0.02 | 96\% |
| 17 | rs9635726 | 38,020,141 | C/T | 0.20 | 1.17 (1.14-1.21) | $1.50 \times 10^{-22}$ | 1.17 (1.14-1.21) | $1.50 \times 10^{-22}$ | 0.54 | 96\% |
| 17 | rs4795397 | 38,023,745 | A/G | 0.48 | 0.86 (0.83-0.9) | $6.09 \times 10^{-15}$ | 0.86 (0.84-0.88) | $2.15 \times 10^{-27}$ | 0.01 | 89\% |
| 17 | rs11557466 | 38,024,626 | C/T | 0.47 | 0.86 (0.83-0.89) | $2.37 \times 10^{-18}$ | 0.85 (0.83-0.88) | $6.18 \times 10^{-32}$ | 0.02 | 95\% |
| 17 | rs11078925 | 38,025,208 | T/C | 0.47 | 0.86 (0.83-0.88) | $2.29 \times 10^{-20}$ | 0.85 (0.83-0.88) | $1.01 \times 10^{-33}$ | 0.03 | 98\% |
| 17 | rs12150079 | 38,025,417 | G/A | 0.32 | 0.88 (0.85-0.91) | $9.80 \times 10^{-13}$ | 0.88 (0.85-0.9) | $1.12 \times 10^{-19}$ | 0.05 | 96\% |
| 17 | rs11557467 | 38,028,634 | G/T | 0.52 | 0.85 (0.82-0.88) | $9.94 \times 10^{-19}$ | 0.85 (0.83-0.87) | $1.39 \times 10^{-29}$ | 0.04 | 88\% |
| 17 | rs10852936 | 38,031,714 | C/T | 0.47 | 0.86 (0.83-0.89) | $1.84 \times 10^{-15}$ | 0.86 (0.83-0.88) | $2.00 \times 10^{-26}$ | 0.02 | 88\% |
| 17 | rs1054609 | 38,033,277 | A/C | 0.47 | 0.85 (0.83-0.88) | $5.07 \times 10^{-20}$ | 0.85 (0.83-0.87) | $8.20 \times 10^{-34}$ | 0.02 | 96\% |
| 17 | rs9907088 | 38,035,116 | G/A | 0.47 | 0.86 (0.83-0.88) | $2.32 \times 10^{-20}$ | 0.85 (0.83-0.88) | $6.85 \times 10^{-34}$ | 0.02 | 98\% |
| 17 | rs12232497 | 38,040,119 | T/C | 0.47 | 0.86 (0.83-0.88) | $2.39 \times 10^{-20}$ | 0.85 (0.83-0.88) | $5.47 \times 10^{-34}$ | 0.02 | 98\% |
| 17 | rs2872507 | 38,040,763 | G/A | 0.47 | 0.85 (0.83-0.88) | $2.76 \times 10^{-20}$ | 0.85 (0.83-0.88) | $5.24 \times 10^{-34}$ | 0.02 | 98\% |
| 17 | rs9901146 | 38,043,343 | G/A | 0.52 | 0.84 (0.82-0.87) | $3.29 \times 10^{-25}$ | 0.84 (0.82-0.86) | $1.87 \times 10^{-40}$ | 0.04 | 98\% |
| 17 | rs12950743 | 38,049,233 | T/C | 0.52 | 0.84 (0.82-0.87) | $2.66 \times 10^{-25}$ | 0.84 (0.82-0.86) | $1.75 \times 10^{-40}$ | 0.04 | 98\% |
| 17 | rs7359623 | 38,049,589 | C/T | 0.51 | 0.84 (0.82-0.87) | $1.16 \times 10^{-23}$ | 0.84 (0.82-0.86) | $2.16 \times 10^{-37}$ | 0.05 | 95\% |
| 17 | rs8067378 | 38,051,348 | A/G | 0.52 | 0.84 (0.82-0.87) | $2.57 \times 10^{-25}$ | 0.84 (0.82-0.86) | $2.78 \times 10^{-40}$ | 0.04 | 98\% |
| 17 | rs8069176 | 38,057,197 | G/A | 0.47 | 0.85 (0.82-0.88) | $1.67 \times 10^{-19}$ | 0.85 (0.83-0.87) | $2.92 \times 10^{-36}$ | 0.01 | 98\% |
| 17 | rs2305480 | 38,062,196 | G/A | 0.46 | 0.85 (0.82-0.88) | $5.62 \times 10^{-19}$ | 0.85 (0.83-0.87) | $3.18 \times 10^{-36}$ | 0.00 | 98\% |
| 17 | rs2305479 | 38,062,217 | C/T | 0.51 | 0.84 (0.81-0.87) | $9.31 \times 10^{-24}$ | 0.84 (0.82-0.86) | $1.00 \times 10^{-42}$ | 0.01 | 98\% |
| 17 | rs11078926 | 38,062,976 | G/A | 0.46 | 0.85 (0.82-0.89) | $6.81 \times 10^{-15}$ | 0.85 (0.83-0.88) | $7.13 \times 10^{-29}$ | 0.00 | 88\% |
| 17 | rs11078927 | 38,064,405 | C/T | 0.46 | 0.85 (0.82-0.88) | $2.31 \times 10^{-19}$ | 0.85 (0.83-0.87) | $4.21 \times 10^{-36}$ | 0.01 | 98\% |
| 17 | rs2290400 | 38,066,240 | T/C | 0.51 | 0.84 (0.81-0.87) | $8.25 \times 10^{-24}$ | 0.84 (0.82-0.86) | $1.03 \times 10^{-42}$ | 0.01 | 98\% |
| 17 | rs1008723 | 38,066,267 | G/T | 0.51 | 0.84 (0.81-0.87) | $3.29 \times 10^{-23}$ | 0.84 (0.82-0.86) | $6.29 \times 10^{-42}$ | 0.01 | 98\% |
| 17 | rs4795400 | 38,067,020 | C/T | 0.47 | 0.85 (0.82-0.88) | $9.86 \times 10^{-19}$ | 0.85 (0.83-0.87) | $1.25 \times 10^{-35}$ | 0.00 | 98\% |
| 17 | rs869402 | 38,068,043 | T/C | 0.49 | 1.19 (1.15-1.23) | $1.35 \times 10^{-21}$ | 1.19 (1.16-1.22) | $8.01 \times 10^{-41}$ | 0.01 | 96\% |
| 17 | rs7216389 | 38,069,949 | C/T | 0.49 | 1.19 (1.15-1.23) | $3.16 \times 10^{-21}$ | 1.2 (1.16-1.23) | $2.51 \times 10^{-42}$ | 0.00 | 98\% |
| 17 | rs9303280 | 38,074,031 | T/C | 0.50 | 1.18 (1.14-1.23) | $6.44 \times 10^{-19}$ | 1.19 (1.16-1.22) | $1.38 \times 10^{-38}$ | 0.00 | 96\% |
| 17 | rs9303281 | 38,074,046 | G/A | 0.48 | 1.19 (1.15-1.23) | $2.49 \times 10^{-21}$ | 1.2 (1.16-1.23) | $2.91 \times 10^{-42}$ | 0.00 | 98\% |
| 17 | rs7219923 | 38,074,518 | C/T | 0.48 | 1.19 (1.15-1.23) | $2.23 \times 10^{-21}$ | 1.2 (1.16-1.23) | $2.81 \times 10^{-42}$ | 0.00 | 98\% |
| 17 | rs7224129 | 38,075,426 | G/A | 0.48 | 1.19 (1.15-1.23) | $1.90 \times 10^{-21}$ | 1.2 (1.17-1.23) | $2.45 \times 10^{-42}$ | 0.00 | 98\% |
| 17 | rs4378650 | 38,080,865 | A/G | 0.51 | 1.19 (1.14-1.24) | $8.37 \times 10^{-17}$ | 1.21 (1.18-1.25) | $2.09 \times 10^{-40}$ | 0.00 | 84\% |
| 17 | rs8076131 | 38,080,912 | G/A | 0.54 | 1.17 (1.13-1.22) | $1.85 \times 10^{-16}$ | 1.18 (1.15-1.21) | $2.44 \times 10^{-34}$ | 0.00 | 96\% |
| 17 | rs12603332 | 38,082,807 | T/C | 0.51 | 1.18 (1.14-1.22) | $3.06 \times 10^{-18}$ | 1.19 (1.16-1.22) | $1.01 \times 10^{-37}$ | 0.00 | 96\% |
| 17 | rs4795405 | 38,088,417 | T/C | 0.55 | 1.17 (1.13-1.21) | $1.66 \times 10^{-16}$ | 1.18 (1.15-1.21) | $1.39 \times 10^{-39}$ | 0.00 | 98\% |
| 17 | rs4794820 | 38,089,344 | A/G | 0.56 | 1.17 (1.12-1.21) | $1.52 \times 10^{-15}$ | 1.18 (1.14-1.21) | $8.05 \times 10^{-32}$ | 0.00 | 95\% |
| 17 | rs7207600 | 38,091,660 | G/A | 0.63 | 1.15 (1.11-1.19) | $1.17 \times 10^{-17}$ | 1.16 (1.13-1.19) | $4.72 \times 10^{-29}$ | 0.07 | 100\% |
| 17 | rs8079416 | 38,092,713 | T/C | 0.44 | 1.17 (1.13-1.2) | $1.45 \times 10^{-22}$ | 1.17 (1.14-1.2) | $1.43 \times 10^{-35}$ | 0.08 | 98\% |


| Chrom | SNP | Position | Allele <br> (R/E) | EAFa | $\begin{aligned} & \text { OR }_{\text {random }} \\ & (95 \% \mathrm{Cl})^{\mathrm{b}} \end{aligned}$ | Prandom ${ }^{\text {c }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{b} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P h e t}^{\text {e }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 17 | rs8065126 | 38,099,035 | T/C | 0.63 | 1.15 (1.11-1.18) | $9.91 \times 10^{-18}$ | 1.16 (1.13-1.19) | $1.79 \times 10^{-27}$ | 0.11 | 98\% |
| 17 | rs4795408 | 38,107,627 | G/A | 0.44 | 1.17 (1.13-1.2) | $2.00 \times 10^{-21}$ | 1.17 (1.15-1.2) | $2.12 \times 10^{-36}$ | 0.05 | 98\% |
| 17 | rs9895948 | 38,108,363 | T/C | 0.64 | 1.14 (1.1-1.18) | $3.69 \times 10^{-14}$ | 1.15 (1.12-1.18) | $1.38 \times 10^{-24}$ | 0.07 | 91\% |
| 17 | rs17609240 | 38,110,689 | T/G | 0.64 | 1.14 (1.1-1.18) | $6.19 \times 10^{-13}$ | 1.15 (1.12-1.19) | $2.86 \times 10^{-24}$ | 0.03 | 91\% |
| 17 | rs1007654 | 38,111,354 | A/G | 0.64 | 1.14 (1.11-1.18) | $9.11 \times 10^{-18}$ | 1.15 (1.12-1.18) | $3.77 \times 10^{-26}$ | 0.15 | 98\% |
| 17 | rs1007655 | 38,111,419 | G/A | 0.63 | 1.14 (1.1-1.18) | $8.70 \times 10^{-14}$ | 1.15 (1.12-1.19) | $1.80 \times 10^{-23}$ | 0.10 | 88\% |
| 17 | rs2313640 | 38,111,845 | C/T | 0.64 | 1.14 (1.11-1.18) | $8.19 \times 10^{-18}$ | 1.15 (1.12-1.18) | $2.96 \times 10^{-26}$ | 0.15 | 98\% |
| 17 | rs7218742 | 38,114,361 | A/G | 0.64 | 1.14 (1.11-1.18) | $4.27 \times 10^{-18}$ | 1.15 (1.12-1.18) | $1.94 \times 10^{-26}$ | 0.16 | 98\% |
| 17 | rs7218321 | 38,114,469 | C/T | 0.64 | 1.14 (1.11-1.18) | $5.22 \times 10^{-16}$ | 1.15 (1.12-1.18) | $3.33 \times 10^{-25}$ | 0.11 | 95\% |
| 17 | rs7219080 | 38,114,516 | A/C | 0.64 | 1.14 (1.11-1.18) | $4.14 \times 10^{-17}$ | 1.15 (1.12-1.18) | $4.90 \times 10^{-26}$ | 0.13 | 98\% |
| 17 | rs6503526 | 38,114,598 | C/T | 0.44 | 1.16 (1.13-1.2) | $5.82 \times 10^{-21}$ | 1.17 (1.14-1.2) | $2.27 \times 10^{-35}$ | 0.05 | 98\% |
| 17 | rs6503527 | 38,114,719 | G/A | 0.64 | 1.14 (1.1-1.18) | $1.35 \times 10^{-13}$ | 1.15 (1.12-1.18) | $4.70 \times 10^{-23}$ | 0.09 | 89\% |
| 17 | rs3902025 | 38,119,254 | G/T | 0.56 | 1.15 (1.11-1.19) | $3.36 \times 10^{-16}$ | 1.16 (1.13-1.19) | $8.34 \times 10^{-31}$ | 0.02 | 98\% |
| 17 | rs3894194 | 38,121,993 | G/A | 0.44 | 1.16 (1.12-1.2) | $8.43 \times 10^{-20}$ | 1.17 (1.14-1.2) | $1.06 \times 10^{-34}$ | 0.03 | 98\% |
| 17 | rs7212938 | 38,122,680 | G/T | 0.55 | 0.86 (0.83-0.88) | $2.65 \times 10^{-20}$ | 0.85 (0.83-0.88) | $7.78 \times 10^{-33}$ | 0.05 | 96\% |
| 17 | rs3859192 | 38,128,648 | C/T | 0.47 | 1.15 (1.11-1.18) | $1.09 \times 10^{-18}$ | 1.15 (1.12-1.18) | $5.38 \times 10^{-28}$ | 0.11 | 96\% |
| 17 | rs8075668 | 38,137,623 | C/T | 0.46 | 0.89 (0.87-0.91) | $1.07 \times 10^{-20}$ | 0.89 (0.87-0.91) | $1.07 \times 10^{-20}$ | 0.66 | 96\% |
| 17 | rs2305481 | 38,138,624 | G/A | 0.37 | 0.89 (0.87-0.91) | $9.27 \times 10^{-20}$ | 0.89 (0.87-0.91) | $9.27 \times 10^{-20}$ | 0.56 | 100\% |
| 17 | rs2305482 | 38,140,927 | A/C | 0.54 | 0.92 (0.9-0.94) | $7.12 \times 10^{-11}$ | 0.92 (0.9-0.94) | $7.12 \times 10^{-11}$ | 0.60 | 100\% |
| 17 | rs11078930 | 38,141,955 | C/T | 0.35 | 0.88 (0.85-0.9) | $1.49 \times 10^{-20}$ | 0.88 (0.85-0.9) | $1.49 \times 10^{-20}$ | 0.65 | 93\% |
| 17 | rs4065321 | 38,143,548 | C/T | 0.54 | 0.92 (0.9-0.94) | $2.17 \times 10^{-11}$ | 0.92 (0.9-0.94) | $2.17 \times 10^{-11}$ | 0.60 | 96\% |
| 17 | rs8066582 | 38,146,929 | T/C | 0.54 | 0.92 (0.9-0.94) | $8.53 \times 10^{-11}$ | 0.92 (0.9-0.94) | $8.53 \times 10^{-11}$ | 0.55 | 100\% |
| 17 | rs11658328 | 38,149,236 | T/C | 0.54 | 0.92 (0.9-0.94) | $7.43 \times 10^{-11}$ | 0.92 (0.9-0.94) | $7.43 \times 10^{-11}$ | 0.62 | 100\% |
| 17 | rs2241245 | 38,151,014 | C/T | 0.54 | 0.92 (0.9-0.94) | $4.72 \times 10^{-11}$ | 0.92 (0.9-0.94) | $4.72 \times 10^{-11}$ | 0.57 | 100\% |
| 17 | rs12453334 | 38,153,473 | C/T | 0.37 | 0.89 (0.87-0.91) | $2.56 \times 10^{-19}$ | 0.89 (0.87-0.91) | $2.56 \times 10^{-19}$ | 0.54 | 100\% |
| 17 | rs4794822 | 38,156,712 | C/T | 0.38 | 1.11 (1.08-1.14) | $5.96 \times 10^{-12}$ | 1.11 (1.08-1.14) | $9.99 \times 10^{-17}$ | 0.13 | 100\% |
| 17 | rs8070454 | 38,160,754 | C/T | 0.38 | 1.11 (1.08-1.14) | $8.97 \times 10^{-12}$ | 1.11 (1.08-1.14) | $1.04 \times 10^{-16}$ | 0.12 | 100\% |
| 17 | rs8078723 | 38,166,879 | T/C | 0.38 | 1.11 (1.08-1.14) | $1.29 \times 10^{-11}$ | 1.11 (1.08-1.14) | $1.03 \times 10^{-16}$ | 0.11 | 100\% |
| 17 | rs2227319 | 38,170,845 | G/A | 0.37 | 0.89 (0.87-0.91) | $6.20 \times 10^{-20}$ | 0.89 (0.87-0.91) | $6.20 \times 10^{-20}$ | 0.58 | 100\% |
| 17 | rs25645 | 38,173,143 | G/A | 0.36 | 0.89 (0.86-0.91) | $5.18 \times 10^{-20}$ | 0.89 (0.86-0.91) | $5.18 \times 10^{-20}$ | 0.50 | 100\% |
| 17 | rs1042658 | 38,173,902 | C/T | 0.38 | 1.11 (1.08-1.14) | $3.45 \times 10^{-12}$ | 1.11 (1.09-1.14) | $6.28 \times 10^{-17}$ | 0.14 | 100\% |
| 17 | rs1045929 | 38,175,426 | C/T | 0.38 | 1.11 (1.07-1.15) | $8.53 \times 10^{-10}$ | 1.11 (1.08-1.14) | $2.76 \times 10^{-14}$ | 0.12 | 89\% |
| 17 | rs12309 | 38,175,462 | C/T | 0.37 | 0.88 (0.86-0.91) | $1.43 \times 10^{-17}$ | 0.88 (0.86-0.91) | $4.83 \times 10^{-18}$ | 0.46 | 89\% |
| 17 | rs709592 | 38,175,553 | C/T | 0.38 | 1.11 (1.08-1.14) | $3.75 \times 10^{-12}$ | 1.11 (1.09-1.14) | $7.06 \times 10^{-17}$ | 0.14 | 100\% |
| 17 | rs2302776 | 38,178,149 | A/G | 0.45 | 0.91 (0.88-0.93) | $4.38 \times 10^{-13}$ | 0.91 (0.88-0.93) | $4.38 \times 10^{-13}$ | 0.83 | 96\% |
| 17 | rs3213762 | 38,178,627 | A/G | 0.38 | 1.11 (1.08-1.15) | $4.78 \times 10^{-12}$ | 1.12 (1.09-1.15) | $3.11 \times 10^{-17}$ | 0.12 | 96\% |
| 17 | rs12451897 | 38,179,275 | G/T | 0.37 | 0.89 (0.87-0.91) | $1.36 \times 10^{-17}$ | 0.89 (0.87-0.91) | $1.36 \times 10^{-17}$ | 0.49 | 91\% |
| 17 | rs2302777 | 38,179,492 | A/G | 0.37 | 0.89 (0.86-0.91) | $1.69 \times 10^{-20}$ | 0.89 (0.86-0.91) | $1.69 \times 10^{-20}$ | 0.64 | 100\% |
| 17 | rs9916158 | 38,182,229 | G/T | 0.37 | 0.89 (0.87-0.91) | $7.10 \times 10^{-18}$ | 0.89 (0.87-0.91) | $7.10 \times 10^{-18}$ | 0.50 | 93\% |
| 17 | rs2302774 | 38,183,090 | G/T | 0.38 | 0.89 (0.86-0.91) | $3.59 \times 10^{-20}$ | 0.89 (0.86-0.91) | $3.59 \times 10^{-20}$ | 0.56 | 98\% |
| 17 | rs7502514 | 38,188,844 | A/G | 0.62 | 0.9 (0.87-0.93) | $8.15 \times 10^{-12}$ | 0.89 (0.87-0.92) | $7.68 \times 10^{-17}$ | 0.13 | 96\% |
| 17 | rs3935280 | 38,189,055 | A/G | 0.62 | 0.9 (0.87-0.93) | $2.00 \times 10^{-9}$ | 0.9 (0.87-0.92) | $4.94 \times 10^{-14}$ | 0.11 | 86\% |
| 17 | rs11078936 | 38,197,914 | T/C | 0.37 | 0.89 (0.87-0.92) | $7.28 \times 10^{-18}$ | 0.89 (0.87-0.92) | $7.28 \times 10^{-18}$ | 0.67 | 98\% |
| 17 | rs8065443 | 38,208,940 | A/G | 0.59 | 0.9 (0.88-0.93) | $1.07 \times 10^{-11}$ | 0.9 (0.88-0.93) | $3.27 \times 10^{-15}$ | 0.19 | 98\% |
| 17 | rs868150 | 38,213,359 | A/G | 0.59 | 0.9 (0.88-0.93) | $6.71 \times 10^{-12}$ | 0.9 (0.88-0.93) | $2.73 \times 10^{-15}$ | 0.20 | 98\% |
| 17 | rs17637472 | 47,461,433 | G/A | 0.39 | 1.08 (1.05-1.11) | $3.28 \times 10^{-9}$ | 1.08 (1.05-1.11) | $3.28 \times 10^{-9}$ | 0.56 | 98\% |
| 17 | rs2671659 | 47,474,529 | G/A | 0.37 | 0.92 (0.9-0.95) | $7.59 \times 10^{-9}$ | 0.92 (0.9-0.95) | $3.70 \times 10-9$ | 0.45 | 96\% |

[^0]Supplementary Table 5. Genetic loci associated with asthma in African ancestry meta-analysis
Results are shown for the lead SNP at each locus reaching $P_{\text {fixed }}<10^{-5}$

| Region ${ }^{\text {a }}$ | Lead SNP | Position | Nearby genes ${ }^{\text {b }}$ | Allele $(R / E)^{c}$ | $E A F^{\text {d }}$ | OR (95\%CI) ${ }^{\text {e }}$ | Pfixed ${ }^{\text {f }}$ | $\mathrm{Phet}^{\text {g }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 1p3.1 | rs10157802 | 74,110,649 | NEGR1,LRRIQ3 | G/A | 0.82 | 0.78 (0.69-0.87) | $7.32 \times 10^{-6}$ | 0.66 |
| 1q23.3 | rs1348135 | 163,776,298 | NUF2,PBX1 | C/T | 0.05 | 0.57 (0.44-0.73) | $8.76 \times 10^{-6}$ | 0.28 |
| 1q42.3 | rs10924970 | 235,362,960 | ARID4B,RBM34,GGPS1 | T/C | 0.61 | 1.25 (1.14-1.37) | $2.87 \times 10^{-6}$ | 0.58 |
| 2p23.3 | rs4268898 | 24,490,413 | ITSN2,C2orf84,NCOA1 | C/T | 0.56 | 0.79 (0.73-0.87) | $1.93 \times 10^{-7}$ | 0.48 |
| 3p24.2 | rs12634582 | 24,282,432 | THRB,NR1D2,RARB | T/C | 0.38 | 1.26 (1.14-1.40) | $9.02 \times 10^{-6}$ | 0.88 |
| 3p22.3 | rs9870718 | 36,239,406 | ARPP21,STAC | C/T | 0.36 | 0.81 (0.74-0.89) | $8.71 \times 10^{-6}$ | 0.75 |
| 7p14.1 | rs10233459 | 42,040,986 | GLI3,INHBA,C7orf25 | A/G | 0.05 | 1.57 (1.30-1.91) | $4.16 \times 10^{-6}$ | 0.51 |
| 11q23.2 | rs4938096 | 114,235,730 | NNMT, C11orf71 | C/T | 0.21 | 1.30 (1.17-1.44) | $1.13 \times 10^{-6}$ | 0.31 |
| 12 24.33 | rs10773588 | 129,511,610 | GLT1D1,TMEM132D | G/T | 0.06 | 0.59 (0.47-0.74) | $4.47 \times 10^{-6}$ | 0.29 |
| 14q12 | rs10141207 | 32,691,785 | ARHGAP5,AKAP6 | G/A | 0.35 | 1.24 (1.13-1.36) | $3.93 \times 10^{-6}$ | 0.40 |
| 20q13.12 | rs16989837 | 43,859,099 | SLP1,SEMG2 | C/T | 0.10 | 1.39 (1.21-1.60) | $3.11 \times 10^{-6}$ | 0.13 |

${ }^{\text {a }}$ Cytogenetic position of the chromosomal region harboring the lead SNP
${ }^{\mathrm{b}}$ Genes coding for proteins; the gene where eventually the lead SNP lies is first indicated followed by the previous gene and next gene
${ }^{c} R=$ reference allele $/ E=$ effect allele.
${ }^{d} E A F=$ Effect allele frequency
${ }^{\text {eO Odds-ratios (ORs) and } 95 \% \text { Confidence Intervals (CI) were computed for the effect allele under a fixed-effects model }}$
${ }^{\mathrm{f}} P_{\text {fixed }}$ is the $P$-value for test of association between SNP and asthma under a fixed-effects model
${ }^{\mathrm{g}} P_{\text {het }}$ is the $P$-value for test of heterogeneity across studies with the use of Cochran's test

Supplementary Table 6. Genetic loci associated with asthma in Japanese meta-analysis Results are shown for the lead SNP at each locus reaching $P_{\text {fixed }}<10^{-5}$

| Region ${ }^{\text {a }}$ | Lead SNP | Position | Nearby genes ${ }^{\text {b }}$ | Allele <br> (R/E) ${ }^{\text {c }}$ | EAF ${ }^{\text {d }}$ | OR (95\%CI) ${ }^{\text {e }}$ | Pfixed ${ }^{\text {f }}$ | Phet ${ }^{\text {g }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 1q32.1 | rs1122396 | 201,138,428 | TMEM9,IGFN1 | G/A | 0.51 | 0.76 (0.68-0.85) | $1.46 \times 10^{-6}$ | 0.83 |
| 7p14.3 | rs10951405 | 34,043,348 | BMPER,BBS9,NPSR1 | C/T | 0.89 | 1.44 (1.24-1.68) | $2.8 \times 10^{-6}$ | 0.28 |
| 11q22.1 | rs3758697 | 100,709,894 | ARHGAP42,CNTN5,TMEM133 | G/A | 0.30 | 1.33 (1.18-1.50) | $2.4 \times 10^{-6}$ | 0.20 |
| 20p13 | rs6084352 | 3,348,407 | C20orf194, SLC4A11, ATRN | A/G | 0.30 | 1.31 (1.16-1.47) | $6.61 \times 10^{-6}$ | 0.77 |

${ }^{\text {a }}$ Cytogenetic position of the chromosomal region harboring the lead SNP
${ }^{\mathrm{b}}$ Genes coding for proteins; the gene where eventually the lead SNP lies is first indicated followed by the previous gene and next gene
${ }^{c} R=$ reference allele $/ E=$ effect allele. ${ }^{d} E A F=E f f e c t ~ a l l e l e ~ f r e q u e n c y ~$
eOdds-ratios (ORs) and 95\% Confidence Intervals (CI) were computed for the effect allele under a fixed-effects model
${ }^{\mathrm{f}} P_{\text {fixed }}$ is the $P$-value for test of association between SNP and asthma under a fixed-effects model
${ }^{\mathrm{g}} P_{\text {het }}$ is the $P$-value for test of heterogeneity across studies using the Cochran's test

Supplementary Table 7. Genetic loci associated with asthma in Latinos
Results are shown for the lead SNP at each locus reaching $P<10^{-5}$

| Region ${ }^{\text {a }}$ | Lead SNP | Position | Nearby genes ${ }^{\text {b }}$ | Allele $(R / E)^{c}$ | EAF ${ }^{\text {d }}$ | OR (95\%CI) ${ }^{\text {e }}$ | P ${ }^{\text {f }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 1p31.1 | rs2352521 | 79,256,105 | IFI44,ELTD1 | C/T | 0.72 | 1.66(1.33-2.06) | $4.95 \times 10^{-6}$ |
| 1q31.3 | rs6694672 | 196,945,789 | CFHR2,CFHR5 | T/G | 0.05 | 2.13(1.52-2.97) | $9.41 \times 10^{-6}$ |
| 5p13.2 | rs12521260 | 37,881,414 | GDNF,EGFLAM | G/T | 0.28 | 1.52(1.27-1.81) | $3.04 \times 10^{-6}$ |
| $8 q 23.3$ | rs12542922 | 117,333,797 | TRPS1,EIF3H | A/G | 0.28 | 0.61(0.49-0.74) | $1.67 \times 10^{-6}$ |
| 8q24.12 | rs2073617 | 119,964,283 | TNFRSF11B,SAMD12,COLEC10 | G/A | 0.37 | 1.46(1.24-1.71) | $4.69 \times 10^{-6}$ |
| 19q13.31 | rs4802207 | 44,600,377 | ZNF224,ZNF284,ZNF225 | C/T | 0.59 | 0.68(0.57-0.81) | $9.51 \times 10^{-6}$ |
| $21 q 21.1$ | rs2825968 | 21,415,693 | TMPRSS15,NCAM2 | T/C | 0.51 | 0.69(0.59-0.81) | $5.8 \times 10^{-6}$ |

${ }^{\text {a }}$ Cytogenetic position of the chromosomal region harboring the lead SNP
${ }^{\mathrm{b}}$ Genes coding for proteins; the gene where eventually the lead SNP lies is first indicated followed by the previous gene and next gene ${ }^{c} R=$ reference allele $/ E=$ effect allele. ${ }^{d} E A F=E f f e c t$ allele frequency
${ }^{\text {e Odds-ratios (ORs) and } 95 \% \text { Confidence Intervals (CI) were computed for the effect allele }}$
${ }^{\mathrm{f}} P$ for test of association between SNP and asthma

Supplementary Table 8. SNPs with $P_{\text {random }} \leq 10^{-8}$ in the multi-ancestry meta-analysis

| Chrom | SNP | Position | Allele <br> (R/E) | $\begin{aligned} & \text { OR }_{\text {random }} \\ & \text { (95\% CI) } \end{aligned}$ | Prandom ${ }^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{CI})^{a} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 | rs1465325 | 102,734,551 | A/G | 0.88 (0.85-0.91) | $5.6 \times 10^{-12}$ | 0.88 (0.85-0.91) | $3.5 \times 10^{-15}$ | 0.20 | 100\% |
| 2 | rs1465324 | 102,735,612 | G/A | 0.89 (0.86-0.93) | $2.6 \times 10^{-10}$ | 0.89 (0.87-0.92) | $2.1 \times 10^{-13}$ | 0.19 | 100\% |
| 2 | rs12328681 | 102,736,867 | G/T | 0.90 (0.87-0.93) | $2.0 \times 10^{-10}$ | 0.89 (0.87-0.92) | $4.3 \times 10^{-13}$ | 0.22 | 100\% |
| 2 | rs10172039 | 102,738,785 | C/A | 0.90 (0.87-0.93) | $8.1 \times 10^{-11}$ | 0.89 (0.87-0.92) | $1.9 \times 10^{-13}$ | 0.24 | 100\% |
| 2 | rs12619383 | 102,741,718 | A/G | 0.88 (0.85-0.92) | $3.0 \times 10^{-11}$ | 0.88 (0.86-0.91) | $6.6 \times 10^{-15}$ | 0.15 | 100\% |
| 2 | rs12620132 | 102,742,115 | A/G | 0.88 (0.85-0.92) | $2.0 \times 10^{-11}$ | 0.88 (0.86-0.91) | $4.3 \times 10^{-15}$ | 0.15 | 100\% |
| 2 | rs13392285 | 102,744,338 | C/A | 0.89 (0.86-0.92) | $5.9 \times 10^{-12}$ | 0.89 (0.86-0.92) | $3.3 \times 10^{-14}$ | 0.29 | 100\% |
| 2 | rs13387400 | 102,750,020 | G/A | 0.88 (0.85-0.91) | $1.2 \times 10^{-11}$ | 0.88 (0.85-0.91) | $2.2 \times 10^{-15}$ | 0.16 | 100\% |
| 2 | rs1558641 | 102,765,865 | G/A | 0.89 (0.85-0.92) | $2.4 \times 10^{-09}$ | 0.88 (0.86-0.91) | $2.9 \times 10^{-14}$ | 0.06 | 100\% |
| 2 | rs1558640 | 102,765,878 | G/A | 0.88 (0.85-0.92) | $3.0 \times 10^{-11}$ | 0.88 (0.85-0.91) | $1.4 \times 10^{-15}$ | 0.10 | 100\% |
| 2 | rs949963 | 102,769,786 | C/T | 0.89 (0.86-0.92) | $2.3 \times 10^{-10}$ | 0.89 (0.86-0.91) | $2.7 \times 10^{-15}$ | 0.05 | 100\% |
| 2 | rs871657 | 102,771,341 | C/T | 0.89 (0.86-0.92) | $5.4 \times 10^{-11}$ | 0.89 (0.86-0.91) | $7.9 \times 10^{-16}$ | 0.08 | 100\% |
| 2 | rs2287048 | 102,773,999 | C/T | 0.89 (0.85-0.92) | $5.5 \times 10^{-10}$ | 0.88 (0.86-0.91) | $1.5 \times 10^{-14}$ | 0.08 | 100\% |
| 2 | rs951192 | 102,785,854 | A/C | 0.92 (0.90-0.95) | $2.4 \times 10^{-09}$ | 0.92 (0.90-0.95) | $1.3 \times 10^{-09}$ | 0.44 | 100\% |
| 2 | rs1030021 | 102,801,478 | A/C | 0.93 (0.90-0.95) | $5.3 \times 10^{-09}$ | 0.93 (0.90-0.95) | $3.9 \times 10^{-09}$ | 0.46 | 100\% |
| 2 | rs10189629 | 102,879,464 | C/A | 0.85 (0.82-0.89) | $7.1 \times 10^{-13}$ | 0.86 (0.83-0.89) | $3.2 \times 10^{-17}$ | 0.06 | 98\% |
| 2 | rs11692065 | 102,883,975 | C/T | 0.85 (0.82-0.89) | $4.3 \times 10^{-13}$ | 0.86 (0.83-0.89) | $2.6 \times 10^{-17}$ | 0.07 | 98\% |
| 2 | rs11674302 | 102,887,128 | T/C | 0.85 (0.82-0.89) | $3.2 \times 10^{-12}$ | 0.86 (0.83-0.89) | $2.4 \times 10^{-17}$ | 0.03 | 98\% |
| 2 | rs11690644 | 102,914,214 | A/G | 0.84 (0.80-0.88) | $2.9 \times 10^{-13}$ | 0.84 (0.81-0.87) | $4.9 \times 10^{-20}$ | 0.05 | 88\% |
| 2 | rs950880 | 102,932,562 | C/A | 1.11 (1.08-1.14) | $2.6 \times 10^{-17}$ | 1.11 (1.09-1.14) | $8.3 \times 10^{-19}$ | 0.41 | 100\% |
| 2 | rs13001325 | 102,939,036 | C/T | 1.11 (1.08-1.14) | $3.6 \times 10^{-17}$ | 1.11 (1.09-1.14) | $7.0 \times 10^{-19}$ | 0.40 | 100\% |
| 2 | rs12479210 | 102,949,161 | C/T | 1.11 (1.09-1.14) | $1.3 \times 10^{-17}$ | 1.11 (1.09-1.14) | $5.1 \times 10^{-19}$ | 0.41 | 100\% |
| 2 | rs13019081 | 102,950,822 | A/C | 1.11 (1.09-1.14) | $1.3 \times 10^{-17}$ | 1.11 (1.09-1.14) | $4.9 \times 10^{-19}$ | 0.41 | 100\% |
| 2 | rs17026974 | 102,952,360 | G/A | 1.10 (1.07-1.13) | $3.5 \times 10^{-13}$ | 1.10 (1.07-1.13) | $3.5 \times 10^{-13}$ | 0.59 | 100\% |
| 2 | rs3771180 | 102,953,617 | G/T | 0.83 (0.80-0.87) | $1.7 \times 10^{-18}$ | 0.84 (0.81-0.86) | $7.6 \times 10^{-26}$ | 0.05 | 100\% |
| 2 | rs13431828 | 102,954,653 | C/T | 0.84 (0.81-0.87) | $1.2 \times 10^{-18}$ | 0.84 (0.81-0.87) | $2.0 \times 10^{-24}$ | 0.10 | 100\% |
| 2 | rs13408661 | 102,955,082 | G/A | 0.83 (0.80-0.87) | $2.1 \times 10^{-18}$ | 0.84 (0.81-0.87) | $2.9 \times 10^{-25}$ | 0.06 | 100\% |
| 2 | rs873022 | 102,955,683 | G/T | 1.10 (1.07-1.13) | $2.6 \times 10^{-13}$ | 1.10 (1.07-1.13) | $2.6 \times 10^{-13}$ | 0.57 | 100\% |
| 2 | rs3771177 | 102,955,860 | G/T | 1.10 (1.07-1.13) | $2.5 \times 10^{-13}$ | 1.10 (1.07-1.13) | $2.5 \times 10^{-13}$ | 0.57 | 100\% |
| 2 | rs10173081 | 102,957,348 | C/T | 0.83 (0.80-0.87) | $1.9 \times 10^{-18}$ | 0.84 (0.81-0.87) | $2.6 \times 10^{-25}$ | 0.06 | 100\% |
| 2 | rs3732129 | 102,957,532 | T/C | 1.10 (1.07-1.13) | $3.2 \times 10^{-13}$ | 1.10 (1.07-1.13) | $3.2 \times 10^{-13}$ | 0.56 | 100\% |
| 2 | rs1420101 | 102,957,716 | C/T | 1.12 (1.09-1.15) | $3.9 \times 10^{-21}$ | 1.12 (1.09-1.15) | $3.9 \times 10^{-21}$ | 0.61 | 100\% |
| 2 | rs12905 | 102,960,007 | G/A | 1.10 (1.07-1.13) | $4.0 \times 10^{-13}$ | 1.10 (1.07-1.13) | $4.0 \times 10^{-13}$ | 0.55 | 100\% |
| 2 | rs12712142 | 102,960,584 | C/A | 1.12 (1.09-1.14) | $1.5 \times 10^{-18}$ | 1.12 (1.09-1.14) | $1.9 \times 10^{-20}$ | 0.38 | 98\% |
| 2 | rs2160203 | 102,960,824 | A/G | 0.88 (0.85-0.91) | $1.4 \times 10^{-16}$ | 0.88 (0.85-0.91) | $1.4 \times 10^{-16}$ | 0.58 | 85\% |
| 2 | rs13017455 | 102,964,742 | C/T | 1.12 (1.09-1.14) | $1.0 \times 10^{-19}$ | 1.12 (1.09-1.14) | $2.7 \times 10^{-20}$ | 0.45 | 100\% |
| 2 | rs12469506 | 102,965,871 | C/T | 1.11 (1.08-1.14) | $2.0 \times 10^{-14}$ | 1.11 (1.08-1.14) | $9.4 \times 10^{-15}$ | 0.45 | 97\% |
| 2 | rs1921622 | 102,966,067 | G/A | 1.09 (1.06-1.11) | $2.4 \times 10^{-12}$ | 1.09 (1.06-1.11) | $1.1 \times 10^{-12}$ | 0.45 | 100\% |
| 2 | rs10208293 | 102,966,310 | G/A | 0.88 (0.85-0.92) | $3.6 \times 10^{-12}$ | 0.88 (0.86-0.91) | $9.8 \times 10^{-21}$ | 0.00 | 100\% |
| 2 | rs10197862 | 102,966,549 | A/G | 0.84 (0.80-0.87) | $4.0 \times 10^{-17}$ | 0.84 (0.81-0.87) | $1.7 \times 10^{-24}$ | 0.04 | 100\% |
| 2 | rs1861245 | 102,966,906 | C/T | 0.90 (0.87-0.92) | $1.1 \times 10^{-14}$ | 0.90 (0.88-0.92) | $3.5 \times 10^{-18}$ | 0.17 | 100\% |
| 2 | rs13424006 | 102,967,236 | T/C | 0.89 (0.87-0.91) | $1.2 \times 10^{-16}$ | 0.89 (0.87-0.91) | $1.7 \times 10^{-21}$ | 0.16 | 100\% |
| 2 | rs6751967 | 102,967,413 | T/C | 0.89 (0.87-0.91) | $1.2 \times 10^{-16}$ | 0.89 (0.87-0.91) | $1.6 \times 10^{-21}$ | 0.16 | 100\% |
| 2 | rs6749114 | 102,967,587 | A/C | 0.89 (0.87-0.92) | $1.1 \times 10^{-16}$ | 0.89 (0.87-0.91) | $2.0 \times 10^{-21}$ | 0.16 | 100\% |
| 2 | rs11123923 | 102,967,844 | C/A | 1.11 (1.08-1.14) | $6.8 \times 10^{-15}$ | 1.11 (1.09-1.14) | $9.0 \times 10^{-19}$ | 0.25 | 100\% |
| 2 | rs4988955 | 102,967,928 | A/G | 0.89 (0.87-0.92) | $7.7 \times 10^{-17}$ | 0.89 (0.87-0.91) | $2.6 \times 10^{-21}$ | 0.18 | 100\% |
| 2 | rs4988956 | 102,968,007 | G/A | 0.89 (0.87-0.91) | $4.4 \times 10^{-17}$ | 0.89 (0.87-0.91) | $1.5 \times 10^{-21}$ | 0.19 | 100\% |
| 2 | rs4988957 | 102,968,075 | T/C | 0.89 (0.87-0.92) | $7.6 \times 10^{-17}$ | 0.89 (0.87-0.91) | $2.0 \times 10^{-21}$ | 0.18 | 100\% |
| 2 | rs10204137 | 102,968,212 | A/G | 0.89 (0.86-0.92) | $9.6 \times 10^{-13}$ | 0.89 (0.87-0.92) | $1.8 \times 10^{-17}$ | 0.11 | 85\% |
| 2 | rs4988958 | 102,968,285 | T/C | 0.89 (0.87-0.92) | $7.5 \times 10^{-17}$ | 0.89 (0.87-0.91) | $1.9 \times 10^{-21}$ | 0.18 | 100\% |
| 2 | rs10192157 | 102,968,356 | C/T | 0.89 (0.87-0.92) | $6.6 \times 10^{-17}$ | 0.89 (0.87-0.91) | $2.3 \times 10^{-21}$ | 0.18 | 100\% |
| 2 | rs10206753 | 102,968,362 | T/C | 0.89 (0.87-0.92) | $1.8 \times 10^{-12}$ | 0.89 (0.87-0.92) | $9.7 \times 10^{-17}$ | 0.14 | 83\% |
| 2 | rs7603730 | 102,974,371 | A/C | 0.89 (0.87-0.91) | $3.9 \times 10^{-17}$ | 0.89 (0.87-0.91) | $7.4 \times 10^{-22}$ | 0.17 | 100\% |
| 2 | rs12998521 | 102,974,417 | G/T | 1.11 (1.08-1.14) | $7.2 \times 10^{-13}$ | 1.11 (1.08-1.14) | $3.1 \times 10^{-17}$ | 0.18 | 98\% |
| 2 | rs10170583 | 102,974,764 | G/A | 0.89 (0.86-0.91) | $1.8 \times 10^{-16}$ | 0.89 (0.87-0.91) | $1.0 \times 10^{-21}$ | 0.14 | 100\% |
| 2 | rs10176664 | 102,976,172 | G/A | 0.89 (0.86-0.91) | $2.8 \times 10^{-17}$ | 0.89 (0.87-0.91) | $2.0 \times 10^{-22}$ | 0.15 | 100\% |
| 2 | rs3755276 | 102,978,459 | C/T | 0.89 (0.86-0.91) | $3.0 \times 10^{-17}$ | 0.89 (0.87-0.91) | $2.1 \times 10^{-22}$ | 0.15 | 100\% |
| 2 | rs2287037 | 102,979,028 | C/T | 1.12 (1.09-1.14) | $7.9 \times 10^{-20}$ | 1.12 (1.09-1.14) | $3.1 \times 10^{-20}$ | 0.46 | 100\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ (95 \% \mathrm{Cl})^{a} \end{array} \end{aligned}$ | Prandom ${ }^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 | rs3771172 | 102,985,812 | C/T | 1.10 (1.08-1.13) | $6.1 \times 10^{-14}$ | 1.10 (1.08-1.13) | $2.5 \times 10^{-14}$ | 0.45 | 100\% |
| 2 | rs3771171 | 102,985,950 | T/C | 1.10 (1.07-1.13) | $4.7 \times 10^{-13}$ | 1.10 (1.08-1.13) | $4.2 \times 10^{-14}$ | 0.39 | 97\% |
| 2 | rs2160202 | 102,986,154 | G/A | 1.10 (1.08-1.13) | $4.8 \times 10^{-13}$ | 1.10 (1.08-1.13) | $4.1 \times 10^{-14}$ | 0.39 | 97\% |
| 2 | rs3771166 | 102,986,222 | G/A | 0.88 (0.86-0.90) | $1.1 \times 10^{-21}$ | 0.88 (0.86-0.90) | $1.1 \times 10^{-23}$ | 0.37 | 95\% |
| 2 | rs1974675 | 102,986,375 | G/A | 0.88 (0.86-0.90) | $1.3 \times 10^{-21}$ | 0.88 (0.86-0.90) | $1.1 \times 10^{-23}$ | 0.37 | 95\% |
| 2 | rs10439410 | 102,990,788 | G/T | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.1 \times 10^{-13}$ | 0.32 | 100\% |
| 2 | rs6758936 | 102,991,369 | G/A | 0.91 (0.89-0.94) | $2.0 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.3 \times 10^{-13}$ | 0.33 | 100\% |
| 2 | rs2041739 | 102,994,333 | C/T | 0.91 (0.89-0.94) | $1.7 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.3 \times 10^{-13}$ | 0.34 | 100\% |
| 2 | rs17027037 | 102,994,884 | A/G | 1.10 (1.08-1.13) | $1.9 \times 10^{-14}$ | 1.10 (1.08-1.13) | $1.9 \times 10^{-14}$ | 0.62 | 100\% |
| 2 | rs2080289 | 102,995,020 | G/A | 1.10 (1.08-1.13) | $2.1 \times 10^{-14}$ | 1.10 (1.08-1.13) | $2.1 \times 10^{-14}$ | 0.63 | 100\% |
| 2 | rs10208196 | 102,996,345 | G/A | 0.92 (0.89-0.94) | $1.6 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.4 \times 10^{-13}$ | 0.35 | 100\% |
| 2 | rs11683700 | 102,996,805 | C/T | 1.10 (1.07-1.13) | $1.6 \times 10^{-12}$ | 1.10 (1.07-1.13) | $1.0 \times 10^{-13}$ | 0.38 | 97\% |
| 2 | rs3213733 | 102,997,884 | C/A | 0.88 (0.85-0.91) | $4.7 \times 10^{-12}$ | 0.88 (0.86-0.91) | $1.6 \times 10^{-15}$ | 0.09 | 100\% |
| 2 | rs3213732 | 102,998,279 | A/G | 0.92 (0.89-0.94) | $1.5 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.3 \times 10^{-13}$ | 0.35 | 100\% |
| 2 | rs6760621 | 102,999,952 | T/C | 0.91 (0.89-0.94) | $3.6 \times 10^{-11}$ | 0.91 (0.89-0.94) | $4.7 \times 10^{-12}$ | 0.37 | 80\% |
| 2 | rs1035130 | 103,001,402 | C/T | 1.10 (1.08-1.13) | $1.6 \times 10^{-14}$ | 1.10 (1.08-1.13) | $1.6 \times 10^{-14}$ | 0.64 | 100\% |
| 2 | rs3771161 | 103,003,961 | C/A | 0.87 (0.84-0.91) | $4.1 \times 10^{-09}$ | 0.88 (0.85-0.91) | $8.6 \times 10^{-13}$ | 0.04 | 82\% |
| 2 | rs6706002 | 103,006,104 | A/G | 0.91 (0.89-0.94) | $1.6 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.0 \times 10^{-13}$ | 0.33 | 100\% |
| 2 | rs4851570 | 103,006,387 | A/G | 1.10 (1.08-1.13) | $1.4 \times 10^{-14}$ | 1.10 (1.08-1.13) | $1.4 \times 10^{-14}$ | 0.58 | 100\% |
| 2 | rs6749014 | 103,006,448 | C/T | 0.92 (0.89-0.94) | $1.4 \times 10^{-10}$ | 0.92 (0.89-0.94) | $7.0 \times 10^{-12}$ | 0.31 | 83\% |
| 2 | rs4851004 | 103,009,537 | C/T | 0.92 (0.89-0.94) | $5.9 \times 10^{-13}$ | 0.92 (0.90-0.94) | $8.7 \times 10^{-14}$ | 0.38 | 100\% |
| 2 | rs3771158 | 103,009,894 | A/G | 0.88 (0.85-0.91) | $8.2 \times 10^{-12}$ | 0.88 (0.86-0.91) | $2.9 \times 10^{-15}$ | 0.08 | 100\% |
| 2 | rs2287034 | 103,010,588 | C/A | 1.10 (1.07-1.13) | $6.0 \times 10^{-14}$ | 1.10 (1.07-1.13) | $6.0 \times 10^{-14}$ | 0.55 | 100\% |
| 2 | rs2287033 | 103,011,237 | T/C | 0.91 (0.89-0.94) | $5.6 \times 10^{-13}$ | 0.92 (0.89-0.94) | $8.2 \times 10^{-14}$ | 0.38 | 100\% |
| 2 | rs1135354 | 103,014,302 | T/G | 1.10 (1.07-1.14) | $1.2 \times 10^{-11}$ | 1.10 (1.07-1.13) | $1.2 \times 10^{-12}$ | 0.39 | 80\% |
| 2 | rs1420094 | 103,015,687 | C/T | 0.92 (0.89-0.94) | $4.3 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.2 \times 10^{-13}$ | 0.41 | 100\% |
| 2 | rs17027087 | 103,015,918 | C/T | 1.10 (1.08-1.13) | $2.8 \times 10^{-14}$ | 1.10 (1.08-1.13) | $2.8 \times 10^{-14}$ | 0.65 | 100\% |
| 2 | rs6710528 | 103,016,142 | C/T | 0.92 (0.89-0.94) | $1.6 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.7 \times 10^{-13}$ | 0.36 | 100\% |
| 2 | rs3732124 | 103,018,052 | C/T | 0.92 (0.89-0.94) | $1.2 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.6 \times 10^{-13}$ | 0.37 | 100\% |
| 2 | rs4851571 | 103,019,000 | C/T | 0.92 (0.89-0.94) | $1.1 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.8 \times 10^{-13}$ | 0.38 | 100\% |
| 2 | rs4851572 | 103,019,031 | G/A | 0.92 (0.89-0.94) | $1.1 \times 10^{-12}$ | 0.92 (0.90-0.94) | $1.7 \times 10^{-13}$ | 0.38 | 100\% |
| 2 | rs10202813 | 103,019,740 | G/T | 0.88 (0.85-0.91) | $1.3 \times 10^{-11}$ | 0.89 (0.86-0.91) | $4.6 \times 10^{-15}$ | 0.07 | 100\% |
| 2 | rs6710034 | 103,023,678 | G/A | 0.92 (0.89-0.94) | $3.7 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.4 \times 10^{-13}$ | 0.43 | 100\% |
| 2 | rs10181785 | 103,025,274 | C/T | 0.88 (0.85-0.91) | $7.0 \times 10^{-12}$ | 0.88 (0.86-0.91) | $3.5 \times 10^{-15}$ | 0.09 | 100\% |
| 2 | rs12712148 | 103,025,547 | G/A | 0.88 (0.85-0.92) | $1.3 \times 10^{-11}$ | 0.89 (0.86-0.91) | $1.4 \times 10^{-14}$ | 0.11 | 100\% |
| 2 | rs11687768 | 103,025,738 | A/G | 0.88 (0.85-0.91) | $5.8 \times 10^{-12}$ | 0.88 (0.86-0.91) | $3.1 \times 10^{-15}$ | 0.10 | 100\% |
| 2 | rs10203558 | 103,027,640 | T/C | 0.92 (0.89-0.94) | $7.7 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.6 \times 10^{-13}$ | 0.40 | 100\% |
| 2 | rs10200952 | 103,027,651 | A/C | 0.91 (0.89-0.94) | $1.8 \times 10^{-10}$ | 0.92 (0.89-0.94) | $2.5 \times 10^{-11}$ | 0.37 | 77\% |
| 2 | rs7586983 | 103,028,066 | C/T | 0.88 (0.85-0.91) | $8.9 \times 10^{-12}$ | 0.88 (0.86-0.91) | $2.9 \times 10^{-15}$ | 0.08 | 100\% |
| 2 | rs1420105 | 103,035,119 | T/C | 0.91 (0.89-0.94) | $2.6 \times 10^{-13}$ | 0.92 (0.89-0.94) | $8.3 \times 10^{-14}$ | 0.42 | 100\% |
| 2 | rs2293223 | 103,035,468 | C/T | 0.88 (0.85-0.91) | $1.2 \times 10^{-11}$ | 0.88 (0.86-0.91) | $3.6 \times 10^{-15}$ | 0.07 | 97\% |
| 2 | rs2293224 | 103,035,779 | T/C | 0.92 (0.89-0.94) | $6.0 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.4 \times 10^{-13}$ | 0.41 | 100\% |
| 2 | rs6743516 | 103,036,335 | A/G | 0.92 (0.89-0.94) | $5.6 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.5 \times 10^{-13}$ | 0.41 | 100\% |
| 2 | rs3771156 | 103,036,677 | C/T | 1.10 (1.07-1.13) | $4.7 \times 10^{-13}$ | 1.10 (1.07-1.13) | $6.1 \times 10^{-14}$ | 0.40 | 100\% |
| 2 | rs1420100 | 103,037,002 | C/A | 0.92 (0.89-0.94) | $5.3 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.6 \times 10^{-13}$ | 0.42 | 100\% |
| 2 | rs3771155 | 103,037,826 | A/G | 0.92 (0.89-0.94) | $4.7 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.1 \times 10^{-13}$ | 0.41 | 100\% |
| 2 | rs10206291 | 103,038,863 | T/C | 0.92 (0.89-0.94) | $5.4 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.6 \times 10^{-13}$ | 0.42 | 100\% |
| 2 | rs885088 | 103,039,044 | A/G | 0.92 (0.89-0.94) | $5.1 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.5 \times 10^{-13}$ | 0.42 | 100\% |
| 2 | rs3771154 | 103,039,360 | C/T | 0.92 (0.89-0.94) | $5.7 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.5 \times 10^{-13}$ | 0.41 | 100\% |
| 2 | rs6759479 | 103,040,047 | A/C | 0.92 (0.89-0.94) | $7.3 \times 10^{-13}$ | 0.92 (0.90-0.94) | $2.2 \times 10^{-13}$ | 0.42 | 100\% |
| 2 | rs887972 | 103,040,945 | G/A | 1.10 (1.07-1.14) | $2.7 \times 10^{-11}$ | 1.10 (1.07-1.14) | $2.7 \times 10^{-11}$ | 0.49 | 82\% |
| 2 | rs887971 | 103,041,167 | T/C | 1.10 (1.07-1.13) | $2.8 \times 10^{-14}$ | 1.10 (1.07-1.13) | $2.8 \times 10^{-14}$ | 0.64 | 100\% |
| 2 | rs3755266 | 103,042,712 | G/A | 0.92 (0.90-0.94) | $4.6 \times 10^{-13}$ | 0.92 (0.90-0.94) | $4.6 \times 10^{-13}$ | 0.50 | 100\% |
| 2 | rs7559845 | 103,046,214 | T/G | 0.92 (0.90-0.94) | $4.1 \times 10^{-13}$ | 0.92 (0.90-0.94) | $4.1 \times 10^{-13}$ | 0.48 | 100\% |
| 2 | rs2310300 | 103,049,074 | A/G | 0.92 (0.90-0.94) | $2.8 \times 10^{-13}$ | 0.92 (0.90-0.94) | $2.8 \times 10^{-13}$ | 0.53 | 100\% |
| 2 | rs10166330 | 103,050,390 | C/T | 0.88 (0.85-0.91) | $9.5 \times 10^{-12}$ | 0.88 (0.86-0.91) | $2.4 \times 10^{-15}$ | 0.07 | 100\% |
| 2 | rs11681718 | 103,051,144 | A/G | 1.10 (1.07-1.13) | $3.6 \times 10^{-11}$ | 1.10 (1.07-1.12) | $1.2 \times 10^{-12}$ | 0.33 | 100\% |
| 2 | rs4851582 | 103,051,558 | T/C | 1.10 (1.07-1.13) | $3.8 \times 10^{-13}$ | 1.10 (1.07-1.13) | $3.8 \times 10^{-13}$ | 0.49 | 100\% |
| 2 | rs3755265 | 103,052,816 | C/A | 0.92 (0.90-0.94) | $7.1 \times 10^{-14}$ | 0.92 (0.90-0.94) | $7.1 \times 10^{-14}$ | 0.49 | 100\% |
| 2 | rs10176820 | 103,054,420 | T/C | 0.89 (0.86-0.92) | $1.4 \times 10^{-11}$ | 0.89 (0.86-0.92) | $5.0 \times 10^{-14}$ | 0.17 | 100\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ (95 \% \mathrm{Cl})^{a} \end{array} \end{aligned}$ | $\mathbf{P r a n d o m}^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 | rs2058659 | 103,054,556 | G/A | 0.92 (0.90-0.94) | $7.5 \times 10^{-14}$ | 0.92 (0.90-0.94) | $7.5 \times 10^{-14}$ | 0.56 | 100\% |
| 2 | rs17027166 | 103,055,420 | G/A | 1.10 (1.07-1.13) | $7.6 \times 10^{-13}$ | 1.10 (1.07-1.13) | $5.8 \times 10^{-14}$ | 0.37 | 100\% |
| 2 | rs13021177 | 103,056,493 | A/G | 0.91 (0.89-0.93) | $1.1 \times 10^{-13}$ | 0.91 (0.89-0.93) | $1.1 \times 10^{-14}$ | 0.36 | 100\% |
| 2 | rs10490204 | 103,056,534 | A/C | 1.10 (1.07-1.13) | $9.7 \times 10^{-12}$ | 1.10 (1.07-1.13) | $5.9 \times 10^{-13}$ | 0.36 | 100\% |
| 2 | rs17027179 | 103,057,159 | C/T | 1.10 (1.07-1.13) | $5.5 \times 10^{-13}$ | 1.10 (1.07-1.13) | $4.1 \times 10^{-14}$ | 0.37 | 100\% |
| 2 | rs2075186 | 103,057,251 | G/T | 0.88 (0.85-0.91) | $1.6 \times 10^{-11}$ | 0.89 (0.86-0.91) | $4.4 \times 10^{-15}$ | 0.07 | 100\% |
| 2 | rs10490203 | 103,059,237 | T/G | 1.10 (1.07-1.13) | $5.2 \times 10^{-13}$ | 1.10 (1.07-1.13) | $3.8 \times 10^{-14}$ | 0.37 | 100\% |
| 2 | rs3771150 | 103,060,851 | G/A | 1.10 (1.07-1.13) | $2.1 \times 10^{-11}$ | 1.10 (1.07-1.12) | $1.0 \times 10^{-12}$ | 0.35 | 100\% |
| 2 | rs11123928 | 103,061,286 | G/A | 1.08 (1.05-1.11) | $7.0 \times 10^{-09}$ | 1.08 (1.05-1.11) | $4.8 \times 10^{-09}$ | 0.46 | 97\% |
| 2 | rs17027230 | 103,079,330 | C/T | 1.10 (1.07-1.13) | $3.1 \times 10^{-11}$ | 1.10 (1.07-1.13) | $5.5 \times 10^{-13}$ | 0.29 | 98\% |
| 2 | rs10210176 | 103,079,516 | C/A | 0.88 (0.85-0.91) | $8.6 \times 10^{-12}$ | 0.88 (0.86-0.91) | $6.0 \times 10^{-15}$ | 0.10 | 97\% |
| 2 | rs10172116 | 103,087,573 | C/T | 0.88 (0.85-0.91) | $2.2 \times 10^{-11}$ | 0.88 (0.86-0.91) | $6.2 \times 10^{-15}$ | 0.07 | 98\% |
| 2 | rs4851011 | 103,089,678 | C/T | 1.09 (1.06-1.13) | $5.6 \times 10^{-10}$ | 1.09 (1.07-1.12) | $4.0 \times 10^{-12}$ | 0.24 | 98\% |
| 2 | rs17027255 | 103,090,127 | C/T | 1.10 (1.07-1.13) | $8.9 \times 10^{-11}$ | 1.10 (1.07-1.12) | $1.0 \times 10^{-12}$ | 0.26 | 98\% |
| 2 | rs17027258 | 103,091,540 | A/G | 1.10 (1.07-1.13) | $3.3 \times 10^{-11}$ | 1.10 (1.07-1.12) | $7.5 \times 10^{-13}$ | 0.30 | 98\% |
| 2 | rs13030642 | 103,091,585 | C/A | 0.89 (0.85-0.92) | $5.7 \times 10^{-11}$ | 0.89 (0.86-0.92) | $4.5 \times 10^{-14}$ | 0.09 | 98\% |
| 2 | rs13018263 | 103,092,270 | T/C | 0.90 (0.88-0.93) | $5.0 \times 10^{-10}$ | 0.91 (0.88-0.93) | $2.9 \times 10^{-11}$ | 0.25 | 98\% |
| 5 | rs12187994 | 109,950,981 | A/C | 1.12 (1.08-1.16) | $8.8 \times 10^{-09}$ | 1.12 (1.08-1.16) | $8.8 \times 10^{-09}$ | 0.73 | 100\% |
| 5 | rs12186893 | 109,952,983 | G/A | 1.12 (1.08-1.16) | $7.9 \times 10^{-09}$ | 1.12 (1.08-1.16) | $7.9 \times 10^{-09}$ | 0.75 | 100\% |
| 5 | rs3985087 | 110,072,656 | C/A | 1.12 (1.08-1.17) | $8.5 \times 10^{-09}$ | 1.12 (1.08-1.17) | $8.5 \times 10^{-09}$ | 0.90 | 98\% |
| 5 | rs17132319 | 110,074,677 | T/G | 1.12 (1.08-1.17) | $4.5 \times 10^{-09}$ | 1.12 (1.08-1.17) | $4.5 \times 10^{-09}$ | 0.86 | 100\% |
| 5 | rs665063 | 110,075,414 | C/A | 0.90 (0.86-0.93) | $4.4 \times 10^{-09}$ | 0.90 (0.86-0.93) | $4.4 \times 10^{-09}$ | 0.73 | 100\% |
| 5 | rs12187004 | 110,076,352 | A/G | 1.13 (1.08-1.17) | $4.5 \times 10^{-09}$ | 1.13 (1.08-1.17) | $4.5 \times 10^{-09}$ | 0.85 | 100\% |
| 5 | rs244411 | 110,078,480 | C/T | 0.89 (0.86-0.92) | $1.3 \times 10^{-09}$ | 0.89 (0.86-0.92) | $1.3 \times 10^{-09}$ | 0.70 | 98\% |
| 5 | rs17446534 | 110,079,450 | T/C | 1.13 (1.09-1.18) | $1.8 \times 10^{-09}$ | 1.13 (1.09-1.18) | $1.8 \times 10^{-09}$ | 0.85 | 100\% |
| 5 | rs244923 | 110,080,363 | T/C | 0.89 (0.85-0.92) | $7.5 \times 10^{-10}$ | 0.89 (0.85-0.92) | $7.5 \times 10^{-10}$ | 0.68 | 98\% |
| 5 | rs381661 | 110,085,098 | C/T | 0.86 (0.82-0.89) | $8.1 \times 10^{-12}$ | 0.86 (0.82-0.89) | $8.1 \times 10^{-12}$ | 0.96 | 97\% |
| 5 | rs11548494 | 110,100,318 | G/A | 1.14 (1.09-1.19) | $9.7 \times 10^{-09}$ | 1.14 (1.09-1.19) | $9.7 \times 10^{-09}$ | 0.56 | 95\% |
| 5 | rs6893213 | 110,198,114 | C/T | 1.15 (1.10-1.20) | $3.9 \times 10^{-11}$ | 1.15 (1.10-1.20) | $3.9 \times 10^{-11}$ | 0.48 | 95\% |
| 5 | rs1837253 | 110,401,872 | T/C | 1.16 (1.12-1.20) | $1.6 \times 10^{-18}$ | 1.16 (1.13-1.19) | $2.0 \times 10^{-25}$ | 0.06 | 97\% |
| 5 | rs10455025 | 110,404,999 | A/C | 1.15 (1.12-1.18) | $9.4 \times 10^{-26}$ | 1.15 (1.12-1.18) | $9.4 \times 10^{-26}$ | 0.57 | 98\% |
| 5 | rs1898671 | 110,408,002 | C/T | 1.14 (1.11-1.17) | $4.0 \times 10^{-25}$ | 1.14 (1.11-1.17) | $4.0 \times 10^{-25}$ | 0.52 | 100\% |
| 5 | rs17624673 | 110,457,158 | C/T | 1.14 (1.10-1.17) | $3.6 \times 10^{-19}$ | 1.14 (1.11-1.17) | $9.5 \times 10^{-23}$ | 0.31 | 98\% |
| 5 | rs1043828 | 110,464,008 | T/C | 1.13 (1.10-1.16) | $2.0 \times 10^{-19}$ | 1.13 (1.10-1.16) | $2.7 \times 10^{-22}$ | 0.34 | 100\% |
| 5 | rs1438673 | 110,467,499 | C/T | 0.90 (0.88-0.93) | $4.2 \times 10^{-13}$ | 0.90 (0.88-0.92) | $6.1 \times 10^{-19}$ | 0.10 | 100\% |
| 5 | rs6594499 | 110,470,137 | C/A | 0.91 (0.89-0.93) | $3.5 \times 10^{-12}$ | 0.91 (0.88-0.93) | $1.2 \times 10^{-16}$ | 0.17 | 97\% |
| 5 | rs2162893 | 110,558,064 | A/G | 0.91 (0.88-0.93) | $2.8 \times 10^{-13}$ | 0.91 (0.88-0.93) | $2.8 \times 10^{-13}$ | 0.69 | 80\% |
| 5 | rs1457115 | 110,567,598 | C/T | 1.08 (1.05-1.10) | $2.5 \times 10^{-10}$ | 1.08 (1.05-1.10) | $2.5 \times 10^{-10}$ | 0.89 | 98\% |
| 5 | rs11950562 | 131,652,529 | A/C | 0.92 (0.90-0.95) | $9.0 \times 10^{-11}$ | 0.92 (0.90-0.95) | $9.0 \times 10^{-11}$ | 0.70 | 97\% |
| 5 | rs10058074 | 131,686,146 | G/A | 0.93 (0.91-0.95) | $2.2 \times 10^{-09}$ | 0.93 (0.91-0.95) | $2.2 \times 10^{-09}$ | 0.69 | 97\% |
| 5 | rs4705938 | 131,694,077 | T/C | 0.93 (0.90-0.95) | $1.4 \times 10^{-09}$ | 0.93 (0.90-0.95) | $4.3 \times 10^{-10}$ | 0.40 | 97\% |
| 5 | rs17622208 | 131,717,050 | G/A | 0.92 (0.90-0.95) | $2.6 \times 10^{-10}$ | 0.92 (0.90-0.95) | $1.5 \times 10^{-10}$ | 0.45 | 97\% |
| 5 | rs4540166 | 131,779,857 | T/C | 0.92 (0.90-0.95) | $9.3 \times 10^{-09}$ | 0.92 (0.90-0.95) | $9.3 \times 10^{-09}$ | 0.54 | 100\% |
| 5 | rs4371745 | 131,779,955 | T/C | 0.92 (0.90-0.95) | $9.6 \times 10^{-09}$ | 0.92 (0.90-0.95) | $9.6 \times 10^{-09}$ | 0.55 | 100\% |
| 5 | rs1023518 | 131,793,772 | G/T | 1.08 (1.05-1.11) | $7.1 \times 10^{-09}$ | 1.08 (1.05-1.11) | $1.6 \times 10^{-09}$ | 0.41 | 100\% |
| 5 | rs3857440 | 131,794,069 | G/A | 1.08 (1.05-1.11) | $2.1 \times 10^{-09}$ | 1.08 (1.06-1.11) | $1.0 \times 10^{-09}$ | 0.45 | 100\% |
| 5 | rs11745587 | 131,796,922 | G/A | 1.08 (1.05-1.10) | $1.2 \times 10^{-09}$ | 1.08 (1.05-1.10) | $1.2 \times 10^{-09}$ | 0.49 | 98\% |
| 5 | rs6894249 | 131,797,547 | A/G | 1.09 (1.06-1.11) | $3.5 \times 10^{-11}$ | 1.09 (1.06-1.11) | $1.4 \times 10^{-11}$ | 0.44 | 97\% |
| 5 | rs17622656 | 131,820,997 | G/A | 0.91 (0.89-0.94) | $2.3 \times 10^{-10}$ | 0.91 (0.89-0.94) | $1.2 \times 10^{-11}$ | 0.33 | 94\% |
| 5 | rs736801 | 131,833,599 | C/T | 0.91 (0.89-0.94) | $2.0 \times 10^{-11}$ | 0.91 (0.89-0.94) | $6.7 \times 10^{-12}$ | 0.43 | 94\% |
| 5 | rs17622991 | 131,932,753 | G/A | 1.11 (1.07-1.15) | $3.4 \times 10^{-09}$ | 1.11 (1.07-1.14) | $3.6 \times 10^{-12}$ | 0.04 | 100\% |
| 5 | rs12187537 | 131,939,904 | T/G | 1.10 (1.07-1.14) | $8.7 \times 10^{-09}$ | 1.10 (1.07-1.13) | $1.1 \times 10^{-10}$ | 0.17 | 100\% |
| 5 | rs3798135 | 131,965,109 | C/T | 1.10 (1.07-1.14) | $1.1 \times 10^{-09}$ | 1.10 (1.07-1.13) | $4.3 \times 10^{-12}$ | 0.12 | 100\% |
| 5 | rs3798134 | 131,965,179 | G/A | 1.10 (1.07-1.14) | $1.7 \times 10^{-09}$ | 1.10 (1.07-1.13) | $5.0 \times 10^{-12}$ | 0.10 | 100\% |
| 5 | rs6596087 | 131,968,609 | G/A | 1.10 (1.07-1.14) | $1.5 \times 10^{-09}$ | 1.10 (1.07-1.13) | $6.8 \times 10^{-12}$ | 0.12 | 100\% |
| 5 | rs12653750 | 131,971,902 | C/T | 1.10 (1.07-1.14) | $8.7 \times 10^{-10}$ | 1.10 (1.07-1.13) | $2.0 \times 10^{-11}$ | 0.22 | 100\% |
| 5 | rs2040704 | 131,973,177 | A/G | 1.11 (1.07-1.15) | $5.5 \times 10^{-09}$ | 1.10 (1.07-1.14) | $3.5 \times 10^{-12}$ | 0.04 | 100\% |
| 5 | rs2074369 | 131,973,663 | T/C | 1.11 (1.07-1.15) | $6.0 \times 10^{-09}$ | 1.11 (1.08-1.14) | $6.3 \times 10^{-12}$ | 0.06 | 91\% |
| 5 | rs2240032 | 131,977,127 | C/T | 1.10 (1.07-1.14) | $2.9 \times 10^{-10}$ | 1.10 (1.07-1.13) | $1.5 \times 10^{-11}$ | 0.28 | 100\% |
| 5 | rs2158177 | 131,984,058 | A/G | 1.13 (1.09-1.17) | $7.8 \times 10^{-13}$ | 1.13 (1.09-1.17) | $7.8 \times 10^{-13}$ | 0.52 | 97\% |


| Chrom | SNP | Position | Allele (R/E) | OR ${ }_{\text {random }}$ (95\% Cl) ${ }^{\text {a }}$ | $\mathbf{P r a n d o m}^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathrm{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 5 | rs3091307 | 131,989,136 | A/G | 1.13 (1.09-1.17) | $2.0 \times 10^{-10}$ | 1.12 (1.09-1.15) | $3.8 \times 10^{-14}$ | 0.03 | 97\% |
| 5 | rs1295686 | 131,995,843 | T/C | 0.89 (0.87-0.92) | $2.8 \times 10^{-15}$ | 0.89 (0.87-0.92) | $2.8 \times 10^{-15}$ | 0.67 | 100\% |
| 5 | rs20541 | 131,995,964 | A/G | 0.89 (0.87-0.92) | $5.0 \times 10^{-16}$ | 0.89 (0.87-0.92) | $5.0 \times 10^{-16}$ | 0.77 | 98\% |
| 5 | rs1295685 | 131,996,445 | A/G | 0.89 (0.86-0.91) | $9.4 \times 10^{-16}$ | 0.89 (0.86-0.91) | $9.4 \times 10^{-16}$ | 0.62 | 98\% |
| 5 | rs848 | 131,996,500 | A/C | 0.89 (0.87-0.92) | $9.0 \times 10^{-15}$ | 0.89 (0.87-0.92) | $9.0 \times 10^{-15}$ | 0.51 | 97\% |
| 5 | rs847 | 131,996,669 | T/C | 0.89 (0.87-0.92) | $5.5 \times 10^{-13}$ | 0.89 (0.87-0.92) | $6.9 \times 10^{-14}$ | 0.37 | 98\% |
| 5 | rs4912622 | 141,490,587 | G/A | 1.09 (1.06-1.12) | $8.5 \times 10^{-09}$ | 1.09 (1.06-1.11) | $7.1 \times 10^{-12}$ | 0.12 | 100\% |
| 5 | rs7705042 | 141,492,419 | C/A | 1.09 (1.06-1.12) | $7.9 \times 10^{-09}$ | 1.09 (1.06-1.12) | $3.1 \times 10^{-12}$ | 0.11 | 100\% |
| 6 | rs3132616 | 30,416,456 | A/G | 1.12 (1.08-1.17) | $6.3 \times 10^{-09}$ | 1.12 (1.08-1.17) | $6.3 \times 10^{-09}$ | 0.68 | 89\% |
| 6 | rs3094024 | 30,495,860 | A/G | 1.12 (1.08-1.17) | $7.1 \times 10^{-09}$ | 1.12 (1.08-1.17) | $7.1 \times 10^{-09}$ | 0.67 | 95\% |
| 6 | rs3130117 | 30,508,956 | G/A | 1.12 (1.08-1.17) | $3.4 \times 10^{-09}$ | 1.12 (1.08-1.17) | $3.4 \times 10^{-09}$ | 0.56 | 97\% |
| 6 | rs3130247 | 30,515,043 | T/C | 1.12 (1.08-1.17) | $6.2 \times 10^{-09}$ | 1.12 (1.08-1.17) | $6.2 \times 10^{-09}$ | 0.58 | 97\% |
| 6 | rs3132610 | 30,544,401 | A/G | 1.13 (1.08-1.17) | $1.2 \times 10^{-09}$ | 1.13 (1.08-1.17) | $1.2 \times 10^{-09}$ | 0.64 | 97\% |
| 6 | rs9262130 | 30,603,519 | G/A | 1.12 (1.08-1.17) | $2.3 \times 10^{-09}$ | 1.12 (1.08-1.17) | $2.3 \times 10^{-09}$ | 0.69 | 97\% |
| 6 | rs9262135 | 30,618,906 | A/G | 1.12 (1.08-1.17) | $5.2 \times 10^{-09}$ | 1.12 (1.08-1.17) | $5.2 \times 10^{-09}$ | 0.74 | 95\% |
| 6 | rs9262143 | 30,652,781 | C/T | 1.12 (1.08-1.17) | $3.0 \times 10^{-09}$ | 1.12 (1.08-1.17) | $3.0 \times 10^{-09}$ | 0.75 | 95\% |
| 6 | rs3095330 | 30,693,633 | A/G | 1.11 (1.07-1.15) | $4.1 \times 10^{-09}$ | 1.11 (1.07-1.15) | $3.1 \times 10^{-09}$ | 0.46 | 79\% |
| 6 | rs1059612 | 30,708,955 | C/T | 1.13 (1.09-1.17) | $3.4 \times 10^{-10}$ | 1.13 (1.09-1.17) | $3.4 \times 10^{-10}$ | 0.61 | 97\% |
| 6 | rs2284174 | 30,713,580 | T/C | 1.10 (1.07-1.14) | $1.4 \times 10^{-09}$ | 1.10 (1.07-1.14) | $1.4 \times 10^{-09}$ | 0.60 | 100\% |
| 6 | rs3129973 | 30,721,143 | C/T | 1.12 (1.08-1.16) | $3.5 \times 10^{-09}$ | 1.12 (1.08-1.16) | $3.5 \times 10^{-09}$ | 0.74 | 97\% |
| 6 | rs3095326 | 30,725,841 | C/T | 1.12 (1.08-1.16) | $5.7 \times 10^{-09}$ | 1.12 (1.08-1.16) | $5.7 \times 10^{-09}$ | 0.76 | 97\% |
| 6 | rs3130665 | 30,735,979 | C/T | 1.12 (1.08-1.16) | $2.1 \times 10^{-09}$ | 1.12 (1.08-1.16) | $2.1 \times 10^{-09}$ | 0.58 | 97\% |
| 6 | rs3095336 | 30,738,446 | G/A | 1.12 (1.08-1.16) | $1.2 \times 10^{-09}$ | 1.12 (1.08-1.16) | $1.2 \times 10^{-09}$ | 0.79 | 95\% |
| 6 | rs3132605 | 30,739,972 | A/C | 1.12 (1.08-1.16) | $1.2 \times 10^{-09}$ | 1.12 (1.08-1.16) | $1.2 \times 10^{-09}$ | 0.77 | 95\% |
| 6 | rs3129978 | 30,746,331 | A/G | 1.12 (1.08-1.17) | $1.3 \times 10^{-09}$ | 1.12 (1.08-1.17) | $1.3 \times 10^{-09}$ | 0.75 | 95\% |
| 6 | rs3132600 | 30,746,367 | C/T | 1.12 (1.08-1.17) | $4.3 \times 10^{-09}$ | 1.12 (1.08-1.17) | $4.3 \times 10^{-09}$ | 0.61 | 92\% |
| 6 | rs3130673 | 30,746,519 | G/T | 1.12 (1.08-1.17) | $1.1 \times 10^{-09}$ | 1.12 (1.08-1.17) | $1.1 \times 10^{-09}$ | 0.75 | 95\% |
| 6 | rs3131044 | 30,758,664 | A/C | 1.12 (1.08-1.17) | $9.5 \times 10^{-10}$ | 1.12 (1.08-1.17) | $9.5 \times 10^{-10}$ | 0.61 | 95\% |
| 6 | rs3131050 | 30,760,025 | T/C | 1.12 (1.08-1.17) | $1.7 \times 10^{-09}$ | 1.12 (1.08-1.17) | $1.7 \times 10^{-09}$ | 0.60 | 94\% |
| 6 | rs9262200 | 30,760,725 | G/A | 1.12 (1.08-1.16) | $1.2 \times 10^{-09}$ | 1.12 (1.08-1.16) | $1.2 \times 10^{-09}$ | 0.61 | 97\% |
| 6 | rs3131055 | 30,761,487 | T/C | 1.12 (1.08-1.16) | $1.4 \times 10^{-09}$ | 1.12 (1.08-1.16) | $1.4 \times 10^{-09}$ | 0.59 | 97\% |
| 6 | rs3129984 | 30,761,572 | C/T | 1.12 (1.08-1.16) | $4.6 \times 10^{-09}$ | 1.12 (1.08-1.16) | $4.0 \times 10^{-09}$ | 0.47 | 97\% |
| 6 | rs3129985 | 30,762,542 | C/T | 1.12 (1.08-1.16) | $1.6 \times 10^{-09}$ | 1.12 (1.08-1.16) | $1.6 \times 10^{-09}$ | 0.58 | 97\% |
| 6 | rs3131060 | 30,763,291 | G/A | 1.12 (1.08-1.16) | $1.9 \times 10^{-09}$ | 1.12 (1.08-1.16) | $1.9 \times 10^{-09}$ | 0.60 | 97\% |
| 6 | rs3130641 | 30,764,081 | C/T | 1.12 (1.08-1.16) | $4.1 \times 10^{-09}$ | 1.12 (1.08-1.16) | $4.1 \times 10^{-09}$ | 0.73 | 94\% |
| 6 | rs1264376 | 30,765,579 | C/A | 1.12 (1.08-1.17) | $1.4 \times 10^{-09}$ | 1.12 (1.08-1.17) | $1.4 \times 10^{-09}$ | 0.78 | 89\% |
| 6 | rs3130955 | 31,054,511 | C/A | 1.08 (1.05-1.11) | $6.6 \times 10^{-09}$ | 1.08 (1.05-1.10) | $3.0 \times 10^{-09}$ | 0.35 | 98\% |
| 6 | rs3130564 | 31,101,674 | C/T | 1.10 (1.07-1.14) | $1.6 \times 10^{-10}$ | 1.10 (1.07-1.14) | $1.6 \times 10^{-10}$ | 0.54 | 95\% |
| 6 | rs2596464 | 31,412,961 | T/C | 1.09 (1.06-1.12) | $3.1 \times 10^{-10}$ | 1.09 (1.07-1.12) | $4.2 \times 10^{-14}$ | 0.15 | 98\% |
| 6 | rs2596472 | 31,428,967 | G/A | 1.10 (1.06-1.13) | $3.4 \times 10^{-10}$ | 1.10 (1.06-1.13) | $3.4 \times 10^{-10}$ | 0.74 | 95\% |
| 6 | rs3099844 | 31,448,976 | C/A | 1.12 (1.08-1.16) | $1.4 \times 10^{-09}$ | 1.12 (1.08-1.16) | $1.4 \times 10^{-09}$ | 0.56 | 97\% |
| 6 | rs2855812 | 31,472,720 | G/T | 1.10 (1.07-1.13) | $8.9 \times 10^{-12}$ | 1.10 (1.07-1.13) | $1.9 \times 10^{-12}$ | 0.39 | 100\% |
| 6 | rs3134950 | 32,127,477 | C/A | 1.07 (1.05-1.10) | $5.0 \times 10^{-09}$ | 1.07 (1.05-1.10) | $5.0 \times 10^{-09}$ | 0.55 | 100\% |
| 6 | rs1061808 | 32,136,547 | T/G | 1.07 (1.05-1.10) | $4.9 \times 10^{-09}$ | 1.07 (1.05-1.10) | $4.9 \times 10^{-09}$ | 0.56 | 98\% |
| 6 | rs9368716 | 32,306,090 | G/A | 1.08 (1.05-1.10) | $1.7 \times 10^{-10}$ | 1.08 (1.05-1.10) | $1.7 \times 10^{-10}$ | 0.71 | 100\% |
| 6 | rs9268301 | 32,319,637 | G/A | 1.10 (1.07-1.12) | $2.9 \times 10^{-11}$ | 1.10 (1.07-1.13) | $1.6 \times 10^{-13}$ | 0.34 | 98\% |
| 6 | rs1265762 | 32,321,115 | C/A | 1.07 (1.05-1.10) | $1.8 \times 10^{-09}$ | 1.07 (1.05-1.10) | $1.8 \times 10^{-09}$ | 0.89 | 100\% |
| 6 | rs1265760 | 32,321,872 | T/C | 1.07 (1.05-1.10) | $2.0 \times 10^{-09}$ | 1.07 (1.05-1.10) | $2.0 \times 10^{-09}$ | 0.89 | 100\% |
| 6 | rs1265758 | 32,323,529 | A/G | 1.08 (1.05-1.10) | $6.1 \times 10^{-10}$ | 1.08 (1.05-1.10) | $6.1 \times 10^{-10}$ | 0.81 | 100\% |
| 6 | rs3129909 | 32,325,710 | G/A | 1.07 (1.05-1.10) | $9.3 \times 10^{-10}$ | 1.07 (1.05-1.10) | $9.3 \times 10^{-10}$ | 0.81 | 100\% |
| 6 | rs2395150 | 32,326,045 | G/A | 1.08 (1.05-1.10) | $4.5 \times 10^{-10}$ | 1.08 (1.05-1.10) | $4.5 \times 10^{-10}$ | 0.82 | 98\% |
| 6 | rs6904608 | 32,327,727 | C/T | 1.08 (1.05-1.10) | $3.9 \times 10^{-10}$ | 1.08 (1.05-1.10) | $3.9 \times 10^{-10}$ | 0.85 | 100\% |
| 6 | rs6904636 | 32,327,781 | C/T | 1.07 (1.05-1.10) | $4.2 \times 10^{-09}$ | 1.07 (1.05-1.10) | $4.2 \times 10^{-09}$ | 0.90 | 100\% |
| 6 | rs2073044 | 32,338,986 | C/T | 1.11 (1.08-1.14) | $1.3 \times 10^{-12}$ | 1.11 (1.08-1.14) | $1.4 \times 10^{-13}$ | 0.42 | 94\% |
| 6 | rs9268403 | 32,341,473 | T/C | 1.13 (1.09-1.16) | $1.9 \times 10^{-14}$ | 1.13 (1.10-1.16) | $1.7 \times 10^{-19}$ | 0.14 | 98\% |
| 6 | rs9268429 | 32,345,052 | A/G | 1.12 (1.09-1.15) | $4.7 \times 10^{-13}$ | 1.12 (1.09-1.15) | $3.5 \times 10^{-18}$ | 0.12 | 98\% |
| 6 | rs1980495 | 32,346,794 | A/C | 1.14 (1.10-1.18) | $1.9 \times 10^{-12}$ | 1.14 (1.11-1.17) | $2.5 \times 10^{-18}$ | 0.09 | 82\% |
| 6 | rs8180664 | 32,347,490 | C/T | 1.13 (1.09-1.16) | $2.2 \times 10^{-14}$ | 1.13 (1.10-1.16) | $4.0 \times 10^{-20}$ | 0.10 | 100\% |
| 6 | rs2395157 | 32,348,145 | A/G | 1.13 (1.09-1.16) | $1.6 \times 10^{-14}$ | 1.13 (1.10-1.16) | $1.5 \times 10^{-20}$ | 0.10 | 100\% |
| 6 | rs17423649 | 32,357,133 | C/T | 0.90 (0.87-0.93) | $3.4 \times 10^{-09}$ | 0.90 (0.87-0.93) | $1.3 \times 10^{-09}$ | 0.40 | 98\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR }_{\text {random }} \\ & \text { (95\% CI)a } \end{aligned}$ | $\mathbf{P r a n d o m}^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 6 | rs9268474 | 32,357,165 | T/C | 1.13 (1.09-1.16) | $6.0 \times 10^{-14}$ | 1.13 (1.10-1.16) | $6.1 \times 10^{-20}$ | 0.09 | 100\% |
| 6 | rs12529049 | 32,357,715 | C/T | 0.90 (0.87-0.93) | $3.1 \times 10^{-09}$ | 0.90 (0.87-0.93) | $1.2 \times 10^{-09}$ | 0.40 | 100\% |
| 6 | rs12525722 | 32,358,163 | A/G | 0.90 (0.87-0.93) | $3.4 \times 10^{-09}$ | 0.90 (0.88-0.93) | $1.4 \times 10^{-09}$ | 0.40 | 100\% |
| 6 | rs17423691 | 32,358,345 | C/A | 0.90 (0.87-0.94) | $8.6 \times 10^{-09}$ | 0.90 (0.88-0.93) | $2.0 \times 10^{-09}$ | 0.35 | 100\% |
| 6 | rs17423698 | 32,358,368 | T/C | 0.90 (0.87-0.94) | $9.3 \times 10^{-09}$ | 0.91 (0.88-0.94) | $2.1 \times 10^{-09}$ | 0.35 | 100\% |
| 6 | rs9268477 | 32,359,121 | G/A | 1.13 (1.09-1.17) | $2.7 \times 10^{-12}$ | 1.13 (1.10-1.16) | $1.4 \times 10^{-17}$ | 0.11 | 83\% |
| 6 | rs9268480 | 32,363,844 | C/T | 1.12 (1.09-1.16) | $6.2 \times 10^{-14}$ | 1.13 (1.10-1.15) | $1.6 \times 10^{-19}$ | 0.11 | 100\% |
| 6 | rs4248166 | 32,366,421 | T/C | 0.91 (0.88-0.94) | $4.7 \times 10^{-10}$ | 0.91 (0.88-0.94) | $2.7 \times 10^{-10}$ | 0.45 | 100\% |
| 6 | rs2294881 | 32,367,604 | T/C | 0.91 (0.89-0.94) | $1.7 \times 10^{-09}$ | 0.91 (0.89-0.94) | $1.7 \times 10^{-09}$ | 0.57 | 100\% |
| 6 | rs2294880 | 32,367,722 | A/G | 1.13 (1.09-1.17) | $1.1 \times 10^{-11}$ | 1.14 (1.10-1.17) | $6.4 \times 10^{-17}$ | 0.12 | 79\% |
| 6 | rs3817966 | 32,367,847 | T/C | 1.13 (1.09-1.17) | $1.3 \times 10^{-11}$ | 1.13 (1.10-1.16) | $1.2 \times 10^{-17}$ | 0.07 | 85\% |
| 6 | rs3817963 | 32,368,087 | T/C | 1.12 (1.09-1.16) | $6.4 \times 10^{-13}$ | 1.12 (1.09-1.15) | $2.7 \times 10^{-19}$ | 0.06 | 100\% |
| 6 | rs3817962 | 32,368,314 | C/A | 1.13 (1.09-1.17) | $3.0 \times 10^{-11}$ | 1.13 (1.10-1.17) | $7.3 \times 10^{-17}$ | 0.10 | 79\% |
| 6 | rs2076525 | 32,370,616 | T/C | 1.12 (1.08-1.15) | $1.0 \times 10^{-12}$ | 1.12 (1.09-1.15) | $8.2 \times 10^{-18}$ | 0.12 | 97\% |
| 6 | rs2076524 | 32,370,684 | A/G | 1.13 (1.09-1.16) | $1.4 \times 10^{-14}$ | 1.13 (1.10-1.16) | $6.1 \times 10^{-20}$ | 0.13 | 100\% |
| 6 | rs3793126 | 32,371,619 | A/G | 1.12 (1.09-1.16) | $1.4 \times 10^{-13}$ | 1.12 (1.09-1.15) | $5.8 \times 10^{-19}$ | 0.09 | 100\% |
| 6 | rs3793127 | 32,371,915 | C/T | 1.13 (1.09-1.16) | $1.2 \times 10^{-13}$ | 1.13 (1.10-1.16) | $3.8 \times 10^{-18}$ | 0.22 | 100\% |
| 6 | rs9268493 | 32,375,330 | G/A | 1.13 (1.09-1.16) | $2.1 \times 10^{-11}$ | 1.13 (1.09-1.16) | $7.5 \times 10^{-17}$ | 0.07 | 80\% |
| 6 | rs9268494 | 32,375,352 | A/C | 1.12 (1.09-1.16) | $1.3 \times 10^{-12}$ | 1.12 (1.09-1.16) | $2.6 \times 10^{-17}$ | 0.14 | 88\% |
| 6 | rs9268497 | 32,375,424 | G/A | 1.13 (1.09-1.17) | $1.9 \times 10^{-11}$ | 1.13 (1.10-1.16) | $8.6 \times 10^{-17}$ | 0.11 | 82\% |
| 6 | rs3763309 | 32,375,973 | C/A | 1.13 (1.10-1.17) | $1.7 \times 10^{-14}$ | 1.13 (1.10-1.17) | $1.1 \times 10^{-18}$ | 0.25 | 100\% |
| 6 | rs3763311 | 32,376,176 | C/T | 1.13 (1.09-1.17) | $6.6 \times 10^{-12}$ | 1.13 (1.10-1.16) | $1.4 \times 10^{-16}$ | 0.14 | 82\% |
| 6 | rs3763312 | 32,376,348 | G/A | 1.13 (1.08-1.17) | $3.2 \times 10^{-09}$ | 1.14 (1.10-1.18) | $1.2 \times 10^{-14}$ | 0.14 | 80\% |
| 6 | rs3763316 | 32,376,746 | C/T | 1.12 (1.09-1.16) | $5.1 \times 10^{-14}$ | 1.12 (1.10-1.15) | $9.6 \times 10^{-19}$ | 0.16 | 97\% |
| 6 | rs9268516 | 32,379,489 | C/T | 1.13 (1.10-1.16) | $3.5 \times 10^{-15}$ | 1.13 (1.10-1.16) | $8.0 \times 10^{-21}$ | 0.12 | 100\% |
| 6 | rs9268542 | 32,384,721 | A/G | 1.08 (1.05-1.11) | $8.9 \times 10^{-09}$ | 1.09 (1.07-1.12) | $7.0 \times 10^{-13}$ | 0.14 | 100\% |
| 6 | rs2395163 | 32,387,809 | T/C | 1.12 (1.08-1.16) | $1.5 \times 10^{-10}$ | 1.13 (1.10-1.16) | $5.8 \times 10^{-17}$ | 0.09 | 97\% |
| 6 | rs4321864 | 32,399,187 | C/A | 1.09 (1.06-1.13) | $2.0 \times 10^{-09}$ | 1.11 (1.08-1.13) | $1.9 \times 10^{-15}$ | 0.08 | 100\% |
| 6 | rs9268614 | 32,402,778 | T/G | 1.12 (1.08-1.15) | $2.1 \times 10^{-10}$ | 1.13 (1.10-1.16) | $1.4 \times 10^{-16}$ | 0.10 | 100\% |
| 6 | rs6926374 | 32,409,305 | A/G | 1.08 (1.05-1.11) | $1.4 \times 10^{-09}$ | 1.09 (1.06-1.11) | $5.0 \times 10^{-12}$ | 0.29 | 97\% |
| 6 | rs2239804 | 32,411,523 | T/C | 1.08 (1.06-1.11) | $1.1 \times 10^{-09}$ | 1.09 (1.07-1.12) | $1.6 \times 10^{-13}$ | 0.19 | 100\% |
| 6 | rs2239803 | 32,411,833 | C/T | 1.08 (1.05-1.11) | $1.4 \times 10^{-09}$ | 1.09 (1.06-1.11) | $2.5 \times 10^{-12}$ | 0.28 | 97\% |
| 6 | rs9268831 | 32,427,748 | C/T | 1.09 (1.06-1.12) | $8.5 \times 10^{-09}$ | 1.09 (1.07-1.12) | $8.4 \times 10^{-14}$ | 0.04 | 100\% |
| 6 | rs9268853 | 32,429,643 | T/C | 1.12 (1.09-1.15) | $6.1 \times 10^{-17}$ | 1.12 (1.09-1.15) | $1.8 \times 10^{-18}$ | 0.39 | 98\% |
| 6 | rs9268858 | 32,429,758 | T/C | 1.12 (1.09-1.15) | $5.5 \times 10^{-17}$ | 1.12 (1.09-1.15) | $1.5 \times 10^{-18}$ | 0.39 | 98\% |
| 6 | rs9268877 | 32,431,147 | A/G | 1.09 (1.07-1.12) | $4.0 \times 10^{-13}$ | 1.09 (1.07-1.12) | $4.0 \times 10^{-13}$ | 0.78 | 98\% |
| 6 | rs9268923 | 32,432,835 | C/T | 1.14 (1.11-1.17) | $1.3 \times 10^{-24}$ | 1.14 (1.11-1.17) | $1.3 \times 10^{-24}$ | 0.62 | 94\% |
| 6 | rs2395185 | 32,433,167 | G/T | 1.12 (1.09-1.15) | $3.6 \times 10^{-17}$ | 1.12 (1.09-1.15) | $1.1 \times 10^{-18}$ | 0.40 | 98\% |
| 6 | rs9268969 | 32,434,349 | C/T | 1.15 (1.12-1.18) | $1.6 \times 10^{-25}$ | 1.15 (1.12-1.18) | $1.6 \times 10^{-25}$ | 0.65 | 94\% |
| 6 | rs9268979 | 32,435,044 | T/C | 1.09 (1.06-1.12) | $2.4 \times 10^{-11}$ | 1.09 (1.06-1.12) | $2.4 \times 10^{-11}$ | 0.81 | 95\% |
| 6 | rs9405040 | 32,439,393 | A/C | 1.12 (1.09-1.15) | $3.8 \times 10^{-17}$ | 1.12 (1.09-1.15) | $1.5 \times 10^{-18}$ | 0.40 | 98\% |
| 6 | rs9286790 | 32,439,828 | G/A | 1.12 (1.09-1.15) | $4.8 \times 10^{-18}$ | 1.12 (1.09-1.15) | $4.8 \times 10^{-18}$ | 0.48 | 95\% |
| 6 | rs9269065 | 32,440,172 | T/G | 1.10 (1.07-1.13) | $8.4 \times 10^{-11}$ | 1.10 (1.07-1.13) | $8.4 \times 10^{-11}$ | 0.50 | 74\% |
| 6 | rs9269069 | 32,440,362 | C/T | 1.10 (1.07-1.12) | $2.8 \times 10^{-13}$ | 1.10 (1.07-1.12) | $2.8 \times 10^{-13}$ | 0.83 | 98\% |
| 6 | rs9269070 | 32,440,451 | G/A | 1.09 (1.06-1.12) | $3.5 \times 10^{-10}$ | 1.09 (1.06-1.12) | $3.5 \times 10^{-10}$ | 0.61 | 76\% |
| 6 | rs9269071 | 32,440,467 | T/C | 1.09 (1.07-1.12) | $1.6 \times 10^{-12}$ | 1.09 (1.07-1.12) | $1.6 \times 10^{-12}$ | 0.82 | 95\% |
| 6 | rs9269080 | 32,440,969 | G/A | 1.09 (1.06-1.11) | $3.9 \times 10^{-11}$ | 1.09 (1.06-1.11) | $3.9 \times 10^{-11}$ | 0.69 | 95\% |
| 6 | rs7755212 | 32,441,408 | C/T | 1.09 (1.06-1.12) | $8.7 \times 10^{-12}$ | 1.09 (1.06-1.12) | $8.7 \times 10^{-12}$ | 0.70 | 94\% |
| 6 | rs7739203 | 32,441,555 | A/G | 1.09 (1.07-1.12) | $1.8 \times 10^{-12}$ | 1.09 (1.07-1.12) | $1.8 \times 10^{-12}$ | 0.83 | 95\% |
| 6 | rs7739357 | 32,441,641 | A/G | 1.10 (1.07-1.12) | $1.8 \times 10^{-12}$ | 1.10 (1.07-1.12) | $1.8 \times 10^{-12}$ | 0.80 | 92\% |
| 6 | rs9405112 | 32,445,600 | G/A | 1.12 (1.09-1.15) | $1.8 \times 10^{-16}$ | 1.12 (1.09-1.15) | $1.3 \times 10^{-18}$ | 0.35 | 98\% |
| 6 | rs6916742 | 32,453,191 | T/C | 1.11 (1.08-1.14) | $2.3 \times 10^{-11}$ | 1.11 (1.08-1.14) | $9.9 \times 10^{-16}$ | 0.14 | 94\% |
| 6 | rs2516049 | 32,570,400 | T/C | 1.11 (1.07-1.14) | $2.0 \times 10^{-10}$ | 1.11 (1.08-1.14) | $3.2 \times 10^{-16}$ | 0.04 | 98\% |
| 6 | rs9272346 | 32,604,372 | G/A | 1.16 (1.12-1.19) | $5.7 \times 10^{-24}$ | 1.16 (1.13-1.18) | $8.2 \times 10^{-32}$ | 0.14 | 97\% |
| 6 | rs9272723 | 32,609,427 | T/C | 1.14 (1.11-1.18) | $2.9 \times 10^{-20}$ | 1.14 (1.11-1.17) | $6.9 \times 10^{-26}$ | 0.15 | 97\% |
| 6 | rs6928482 | 32,626,249 | T/C | 1.11 (1.08-1.15) | $1.8 \times 10^{-12}$ | 1.11 (1.09-1.14) | $4.8 \times 10^{-19}$ | 0.03 | 98\% |
| 6 | rs9273363 | 32,626,272 | C/A | 1.10 (1.07-1.14) | $2.7 \times 10^{-12}$ | 1.10 (1.07-1.14) | $2.7 \times 10^{-12}$ | 0.99 | 97\% |
| 6 | rs6906021 | 32,626,311 | T/C | 1.10 (1.06-1.13) | $8.4 \times 10^{-10}$ | 1.10 (1.07-1.12) | $3.9 \times 10^{-14}$ | 0.05 | 97\% |
| 6 | rs9275141 | 32,651,117 | T/G | 1.11 (1.07-1.14) | $1.8 \times 10^{-10}$ | 1.11 (1.08-1.13) | $1.2 \times 10^{-17}$ | 0.01 | 100\% |
| 6 | rs2858330 | 32,658,715 | T/C | 1.10 (1.07-1.14) | $1.1 \times 10^{-10}$ | 1.11 (1.08-1.13) | $7.3 \times 10^{-18}$ | 0.01 | 100\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ (95 \% \mathrm{Cl})^{a} \end{array} \end{aligned}$ | Prandom ${ }^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathbf{P f i x e d}^{\text {c }}$ | $P_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 6 | rs16898264 | 32,677,152 | G/A | 1.09 (1.06-1.11) | $1.3 \times 10^{-09}$ | 1.09 (1.06-1.11) | $1.3 \times 10^{-09}$ | 0.48 | 83\% |
| 6 | rs10947332 | 32,677,440 | G/A | 0.89 (0.85-0.92) | $4.5 \times 10^{-09}$ | 0.89 (0.86-0.93) | $6.6 \times 10^{-10}$ | 0.24 | 98\% |
| 6 | rs3104405 | 32,682,308 | C/A | 1.10 (1.07-1.13) | $1.3 \times 10^{-12}$ | 1.10 (1.07-1.13) | $2.4 \times 10^{-13}$ | 0.41 | 94\% |
| 6 | rs2239701 | 32,805,049 | T/C | 1.08 (1.06-1.11) | $2.3 \times 10^{-10}$ | 1.08 (1.06-1.11) | $2.3 \times 10^{-10}$ | 0.88 | 95\% |
| 6 | rs6454802 | 90,814,199 | C/T | 0.93 (0.90-0.95) | $7.4 \times 10^{-09}$ | 0.93 (0.90-0.95) | $7.4 \times 10^{-09}$ | 0.72 | 88\% |
| 6 | rs11753332 | 90,819,153 | G/A | 0.92 (0.90-0.95) | $7.2 \times 10^{-10}$ | 0.92 (0.90-0.95) | $7.2 \times 10^{-10}$ | 0.81 | 95\% |
| 6 | rs12194007 | 90,823,159 | G/T | 0.92 (0.90-0.95) | $2.8 \times 10^{-09}$ | 0.92 (0.90-0.95) | $2.8 \times 10^{-09}$ | 0.72 | 91\% |
| 6 | rs12199079 | 90,852,258 | T/G | 0.92 (0.90-0.94) | $1.3 \times 10^{-10}$ | 0.92 (0.90-0.94) | $1.3 \times 10^{-10}$ | 0.83 | 97\% |
| 6 | rs1010473 | 90,856,878 | G/T | 0.92 (0.89-0.94) | $1.1 \times 10^{-09}$ | 0.92 (0.89-0.94) | $1.1 \times 10^{-09}$ | 0.89 | 85\% |
| 6 | rs1010474 | 90,857,028 | T/C | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.82 | 97\% |
| 6 | rs17711850 | 90,864,870 | T/C | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.79 | 97\% |
| 6 | rs10455168 | 90,883,525 | T/C | 0.92 (0.89-0.94) | $1.6 \times 10^{-11}$ | 0.92 (0.89-0.94) | $1.6 \times 10^{-11}$ | 0.88 | 97\% |
| 6 | rs10806423 | 90,886,824 | C/T | 0.92 (0.89-0.94) | $1.5 \times 10^{-11}$ | 0.92 (0.89-0.94) | $1.5 \times 10^{-11}$ | 0.88 | 97\% |
| 6 | rs11757155 | 90,941,240 | C/T | 0.91 (0.89-0.94) | $4.0 \times 10^{-12}$ | 0.91 (0.89-0.94) | $4.0 \times 10^{-12}$ | 0.63 | 100\% |
| 6 | rs2021716 | 90,941,289 | C/T | 0.91 (0.89-0.94) | $8.6 \times 10^{-12}$ | 0.91 (0.89-0.94) | $8.6 \times 10^{-12}$ | 0.59 | 97\% |
| 6 | rs17585295 | 90,944,831 | C/T | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.67 | 100\% |
| 6 | rs4707609 | 90,946,479 | T/C | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.92 (0.90-0.94) | $1.1 \times 10^{-10}$ | 0.68 | 100\% |
| 6 | rs2875584 | 90,950,628 | C/T | 0.92 (0.89-0.94) | $8.7 \times 10^{-12}$ | 0.92 (0.89-0.94) | $8.7 \times 10^{-12}$ | 0.62 | 100\% |
| 6 | rs17513531 | 90,951,239 | C/T | 0.92 (0.89-0.94) | $7.9 \times 10^{-12}$ | 0.92 (0.89-0.94) | $7.9 \times 10^{-12}$ | 0.61 | 100\% |
| 6 | rs905670 | 90,958,502 | G/A | 0.91 (0.89-0.94) | $3.2 \times 10^{-12}$ | 0.91 (0.89-0.94) | $3.2 \times 10^{-12}$ | 0.71 | 100\% |
| 6 | rs1847472 | 90,973,159 | C/A | 0.92 (0.89-0.94) | $1.4 \times 10^{-11}$ | 0.92 (0.89-0.94) | $1.4 \times 10^{-11}$ | 0.74 | 98\% |
| 6 | rs943689 | 90,984,035 | C/T | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | 0.76 | 100\% |
| 6 | rs2325291 | 90,986,686 | G/A | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | 0.80 | 100\% |
| 6 | rs2325292 | 90,986,749 | T/C | 0.91 (0.89-0.94) | $1.5 \times 10^{-12}$ | 0.91 (0.89-0.94) | $1.5 \times 10^{-12}$ | 0.81 | 100\% |
| 6 | rs2174281 | 90,987,872 | T/C | 0.92 (0.90-0.95) | $9.1 \times 10^{-11}$ | 0.92 (0.90-0.95) | $9.1 \times 10^{-11}$ | 0.62 | 100\% |
| 6 | rs4142967 | 90,996,349 | C/T | 0.92 (0.90-0.95) | $7.0 \times 10^{-11}$ | 0.92 (0.90-0.95) | $7.0 \times 10^{-11}$ | 0.56 | 100\% |
| 6 | rs12212193 | 90,996,769 | A/G | 0.92 (0.90-0.95) | $6.7 \times 10^{-11}$ | 0.92 (0.90-0.95) | $6.7 \times 10^{-11}$ | 0.58 | 100\% |
| 6 | rs1504215 | 91,006,227 | G/A | 0.91 (0.89-0.94) | $1.5 \times 10^{-12}$ | 0.91 (0.89-0.94) | $1.5 \times 10^{-12}$ | 0.89 | 97\% |
| 6 | rs6925032 | 91,008,027 | C/A | 0.91 (0.89-0.94) | $1.6 \times 10^{-12}$ | 0.91 (0.89-0.94) | $1.6 \times 10^{-12}$ | 0.89 | 97\% |
| 6 | rs1321859 | 91,011,673 | C/T | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | 0.91 (0.89-0.94) | $2.2 \times 10^{-12}$ | 0.87 | 97\% |
| 8 | rs13275219 | 81,259,826 | T/C | 0.93 (0.91-0.95) | $1.2 \times 10^{-09}$ | 0.93 (0.91-0.95) | $1.2 \times 10^{-09}$ | 0.58 | 100\% |
| 8 | rs4739735 | 81,259,877 | C/T | 0.93 (0.91-0.95) | $1.2 \times 10^{-09}$ | 0.93 (0.91-0.95) | $7.5 \times 10^{-10}$ | 0.45 | 100\% |
| 8 | rs3913969 | 81,261,064 | C/T | 0.93 (0.91-0.95) | $1.6 \times 10^{-09}$ | 0.93 (0.91-0.95) | $1.6 \times 10^{-09}$ | 0.49 | 100\% |
| 8 | rs7462675 | 81,263,962 | C/A | 0.93 (0.91-0.95) | $1.0 \times 10^{-09}$ | 0.93 (0.91-0.95) | $1.0 \times 10^{-09}$ | 0.51 | 100\% |
| 8 | rs1911713 | 81,266,044 | G/A | 0.92 (0.90-0.95) | $2.3 \times 10^{-10}$ | 0.92 (0.90-0.95) | $1.4 \times 10^{-10}$ | 0.45 | 98\% |
| 8 | rs6992476 | 81,267,706 | A/G | 0.93 (0.91-0.95) | $9.9 \times 10^{-10}$ | 0.93 (0.91-0.95) | $9.9 \times 10^{-10}$ | 0.51 | 100\% |
| 8 | rs7012968 | 81,267,808 | T/C | 0.93 (0.90-0.95) | $8.9 \times 10^{-10}$ | 0.93 (0.90-0.95) | $1.5 \times 10^{-10}$ | 0.37 | 100\% |
| 8 | rs6473222 | 81,267,937 | A/G | 0.93 (0.90-0.95) | $7.7 \times 10^{-10}$ | 0.93 (0.90-0.95) | $1.3 \times 10^{-10}$ | 0.37 | 100\% |
| 8 | rs6473223 | 81,268,155 | T/C | 0.92 (0.90-0.95) | $1.3 \times 10^{-10}$ | 0.92 (0.90-0.95) | $1.3 \times 10^{-10}$ | 0.49 | 100\% |
| 8 | rs6987042 | 81,273,883 | A/G | 0.93 (0.90-0.95) | $5.5 \times 10^{-10}$ | 0.93 (0.90-0.95) | $1.2 \times 10^{-10}$ | 0.39 | 100\% |
| 8 | rs11786704 | 81,275,860 | A/C | 0.93 (0.90-0.95) | $9.1 \times 10^{-10}$ | 0.93 (0.90-0.95) | $2.1 \times 10^{-10}$ | 0.38 | 100\% |
| 8 | rs13275449 | 81,276,113 | G/A | 0.93 (0.90-0.95) | $6.8 \times 10^{-10}$ | 0.93 (0.90-0.95) | $6.8 \times 10^{-10}$ | 0.60 | 97\% |
| 8 | rs12543811 | 81,278,885 | G/A | 0.92 (0.90-0.95) | $1.1 \times 10^{-10}$ | 0.92 (0.90-0.95) | $1.1 \times 10^{-10}$ | 0.54 | 100\% |
| 8 | rs13270496 | 81,280,666 | A/C | 0.93 (0.91-0.95) | $1.3 \times 10^{-09}$ | 0.93 (0.91-0.95) | $1.3 \times 10^{-09}$ | 0.50 | 97\% |
| 8 | rs6473225 | 81,281,007 | G/A | 0.93 (0.90-0.95) | $2.8 \times 10^{-10}$ | 0.93 (0.90-0.95) | $2.8 \times 10^{-10}$ | 0.49 | 100\% |
| 8 | rs7837153 | 81,283,376 | A/C | 0.93 (0.90-0.95) | $5.1 \times 10^{-10}$ | 0.93 (0.90-0.95) | $5.1 \times 10^{-10}$ | 0.64 | 97\% |
| 9 | rs380568 | 6,055,531 | T/C | 0.88 (0.85-0.92) | $4.3 \times 10^{-09}$ | 0.90 (0.88-0.93) | $1.4 \times 10^{-10}$ | 0.01 | 100\% |
| 9 | rs343490 | 6,064,575 | A/G | 0.88 (0.85-0.92) | $2.4 \times 10^{-09}$ | 0.90 (0.87-0.93) | $6.9 \times 10^{-11}$ | 0.01 | 100\% |
| 9 | rs343476 | 6,072,597 | T/C | 1.14 (1.09-1.20) | $1.8 \times 10^{-09}$ | 1.12 (1.08-1.15) | $3.7 \times 10^{-11}$ | 0.01 | 98\% |
| 9 | rs189348 | 6,073,194 | T/C | 1.14 (1.09-1.20) | $3.3 \times 10^{-09}$ | 1.12 (1.08-1.15) | $8.8 \times 10^{-11}$ | 0.01 | 98\% |
| 9 | rs378952 | 6,078,146 | C/T | 1.13 (1.09-1.18) | $8.7 \times 10^{-10}$ | 1.11 (1.08-1.15) | $1.6 \times 10^{-11}$ | 0.03 | 98\% |
| 9 | rs371454 | 6,078,614 | C/T | 1.13 (1.09-1.18) | $9.8 \times 10^{-10}$ | 1.11 (1.08-1.15) | $1.7 \times 10^{-11}$ | 0.02 | 98\% |
| 9 | rs340921 | 6,090,160 | G/T | 1.12 (1.08-1.17) | $3.0 \times 10^{-09}$ | 1.11 (1.08-1.14) | $4.2 \times 10^{-11}$ | 0.03 | 98\% |
| 9 | rs531759 | 6,091,996 | C/T | 1.13 (1.08-1.17) | $2.0 \times 10^{-09}$ | 1.11 (1.08-1.15) | $1.9 \times 10^{-11}$ | 0.02 | 98\% |
| 9 | rs639247 | 6,092,089 | T/G | 1.13 (1.08-1.17) | $2.0 \times 10^{-09}$ | 1.11 (1.08-1.15) | $2.0 \times 10^{-11}$ | 0.02 | 98\% |
| 9 | rs9408638 | 6,096,931 | A/G | 1.13 (1.08-1.17) | $2.0 \times 10^{-09}$ | 1.11 (1.08-1.14) | $3.3 \times 10^{-11}$ | 0.03 | 98\% |
| 9 | rs437389 | 6,099,531 | T/C | 1.13 (1.09-1.18) | $8.5 \times 10^{-10}$ | 1.11 (1.08-1.15) | $8.9 \times 10^{-12}$ | 0.03 | 98\% |
| 9 | rs340906 | 6,106,086 | T/C | 1.13 (1.09-1.17) | $1.7 \times 10^{-09}$ | 1.11 (1.08-1.15) | $2.2 \times 10^{-11}$ | 0.02 | 98\% |
| 9 | rs340905 | 6,106,169 | A/C | 1.13 (1.09-1.17) | $1.4 \times 10^{-09}$ | 1.11 (1.08-1.15) | $1.5 \times 10^{-11}$ | 0.02 | 98\% |
| 9 | rs340904 | 6,106,779 | C/A | 1.13 (1.09-1.18) | $1.7 \times 10^{-09}$ | 1.11 (1.08-1.15) | $1.6 \times 10^{-11}$ | 0.02 | 98\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ \text { ( } 95 \% \mathrm{Cl})^{a} \end{array} \end{aligned}$ | Prandom ${ }^{\text {b }}$ | $\begin{gathered} \text { OR }_{\text {fixed }} \\ (95 \% \mathrm{Cl})^{a} \end{gathered}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathrm{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 9 | rs974936 | 6,111,703 | A/C | 1.13 (1.09-1.17) | $1.1 \times 10^{-09}$ | 1.11 (1.08-1.15) | $1.2 \times 10^{-11}$ | 0.03 | 98\% |
| 9 | rs441616 | 6,113,940 | T/C | 1.13 (1.09-1.18) | $1.4 \times 10^{-09}$ | 1.11 (1.08-1.15) | $1.0 \times 10^{-11}$ | 0.02 | 98\% |
| 9 | rs1556470 | 6,115,538 | C/T | 1.13 (1.09-1.18) | $1.4 \times 10^{-09}$ | 1.11 (1.08-1.15) | $8.6 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs374672 | 6,119,038 | C/T | 1.13 (1.09-1.18) | $1.2 \times 10^{-09}$ | 1.11 (1.08-1.15) | $7.0 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs443175 | 6,123,556 | T/C | 1.13 (1.09-1.18) | $8.0 \times 10^{-10}$ | 1.11 (1.08-1.15) | $3.8 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs1755531 | 6,124,250 | T/G | 1.13 (1.09-1.18) | $1.2 \times 10^{-09}$ | 1.11 (1.08-1.15) | $6.8 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs1970089 | 6,124,359 | T/C | 1.13 (1.09-1.18) | $7.8 \times 10^{-10}$ | 1.12 (1.08-1.15) | $3.5 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs1537285 | 6,124,584 | T/G | 1.13 (1.09-1.18) | $1.2 \times 10^{-09}$ | 1.11 (1.08-1.15) | $6.2 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs1332292 | 6,124,862 | T/C | 1.13 (1.09-1.18) | $1.2 \times 10^{-09}$ | 1.11 (1.08-1.15) | $6.7 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs7039066 | 6,125,539 | T/C | 1.13 (1.09-1.18) | $1.2 \times 10^{-09}$ | 1.11 (1.08-1.15) | $6.9 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs340915 | 6,126,588 | A/G | 1.13 (1.09-1.18) | $1.2 \times 10^{-09}$ | 1.11 (1.08-1.15) | $6.7 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs340914 | 6,126,799 | A/G | 1.13 (1.09-1.18) | $1.1 \times 10^{-09}$ | 1.11 (1.08-1.15) | $6.5 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs340913 | 6,127,330 | T/C | 1.13 (1.09-1.18) | $1.6 \times 10^{-09}$ | 1.11 (1.08-1.15) | $8.3 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs340912 | 6,127,851 | A/G | 1.13 (1.09-1.18) | $8.8 \times 10^{-10}$ | 1.11 (1.08-1.15) | $4.1 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs340908 | 6,128,897 | T/C | 1.13 (1.09-1.18) | $8.8 \times 10^{-10}$ | 1.11 (1.08-1.15) | $5.2 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs340907 | 6,129,637 | A/C | 1.13 (1.09-1.18) | $7.5 \times 10^{-10}$ | 1.11 (1.08-1.15) | $5.4 \times 10^{-12}$ | 0.03 | 98\% |
| 9 | rs1888906 | 6,131,460 | G/A | 1.13 (1.09-1.18) | $6.8 \times 10^{-10}$ | 1.12 (1.08-1.15) | $3.7 \times 10^{-12}$ | 0.02 | 98\% |
| 9 | rs376690 | 6,134,926 | C/T | 1.13 (1.08-1.17) | $1.8 \times 10^{-09}$ | 1.11 (1.08-1.15) | $1.5 \times 10^{-11}$ | 0.03 | 98\% |
| 9 | rs413382 | 6,142,948 | C/A | 1.13 (1.09-1.18) | $3.8 \times 10^{-10}$ | 1.12 (1.08-1.15) | $2.3 \times 10^{-12}$ | 0.05 | 95\% |
| 9 | rs386880 | 6,144,333 | C/T | 1.09 (1.06-1.12) | $3.5 \times 10^{-09}$ | 1.09 (1.06-1.11) | $2.2 \times 10^{-10}$ | 0.23 | 98\% |
| 9 | rs369756 | 6,146,441 | G/T | 1.11 (1.08-1.15) | $1.1 \times 10^{-09}$ | 1.11 (1.08-1.14) | $2.3 \times 10^{-12}$ | 0.15 | 95\% |
| 9 | rs450108 | 6,153,485 | T/C | 1.10 (1.06-1.14) | $8.4 \times 10^{-09}$ | 1.10 (1.07-1.12) | $1.5 \times 10^{-11}$ | 0.11 | 82\% |
| 9 | rs1116795 | 6,155,226 | G/T | 1.09 (1.06-1.12) | $8.1 \times 10^{-09}$ | 1.09 (1.06-1.11) | $1.0 \times 10^{-11}$ | 0.08 | 98\% |
| 9 | rs10124250 | 6,161,686 | C/T | 1.09 (1.06-1.12) | $6.1 \times 10^{-09}$ | 1.09 (1.06-1.11) | $2.6 \times 10^{-12}$ | 0.05 | 98\% |
| 9 | rs10119713 | 6,163,823 | G/A | 1.09 (1.06-1.12) | $3.8 \times 10^{-09}$ | 1.09 (1.06-1.11) | $3.2 \times 10^{-12}$ | 0.07 | 98\% |
| 9 | rs7032572 | 6,172,380 | A/G | 1.18 (1.13-1.23) | $1.4 \times 10^{-16}$ | 1.18 (1.14-1.22) | $1.3 \times 10^{-24}$ | 0.06 | 97\% |
| 9 | rs1412426 | 6,188,652 | A/C | 0.90 (0.87-0.93) | $4.2 \times 10^{-10}$ | 0.90 (0.88-0.92) | $5.9 \times 10^{-18}$ | 0.00 | 100\% |
| 9 | rs1412425 | 6,188,740 | A/C | 0.90 (0.87-0.93) | $2.0 \times 10^{-10}$ | 0.90 (0.87-0.92) | $2.2 \times 10^{-18}$ | 0.00 | 100\% |
| 9 | rs1342326 | 6,190,076 | A/C | 1.17 (1.12-1.21) | $8.6 \times 10^{-14}$ | 1.17 (1.14-1.21) | $2.9 \times 10^{-24}$ | 0.01 | 97\% |
| 9 | rs2095044 | 6,192,796 | T/C | 0.86 (0.83-0.89) | $1.3 \times 10^{-17}$ | 0.87 (0.84-0.89) | $2.1 \times 10^{-27}$ | 0.01 | 100\% |
| 9 | rs2381416 | 6,193,455 | C/A | 0.86 (0.83-0.89) | $1.3 \times 10^{-17}$ | 0.86 (0.84-0.89) | $4.8 \times 10^{-28}$ | 0.01 | 100\% |
| 9 | rs10815370 | 6,194,831 | C/A | 0.90 (0.86-0.93) | $4.6 \times 10^{-10}$ | 0.90 (0.87-0.92) | $4.7 \times 10^{-18}$ | 0.00 | 98\% |
| 9 | rs1888909 | 6,197,392 | T/C | 0.86 (0.83-0.89) | $2.4 \times 10^{-18}$ | 0.86 (0.84-0.89) | $1.3 \times 10^{-28}$ | 0.02 | 100\% |
| 9 | rs992969 | 6,209,697 | A/G | 0.86 (0.83-0.88) | $7.2 \times 10^{-20}$ | 0.86 (0.84-0.88) | $3.3 \times 10^{-30}$ | 0.02 | 100\% |
| 9 | rs3939286 | 6,210,099 | T/C | 0.86 (0.83-0.89) | $1.9 \times 10^{-17}$ | 0.86 (0.84-0.89) | $2.4 \times 10^{-28}$ | 0.01 | 100\% |
| 9 | rs928413 | 6,213,387 | G/A | 0.86 (0.83-0.89) | $3.5 \times 10^{-16}$ | 0.86 (0.84-0.89) | $2.1 \times 10^{-27}$ | 0.00 | 98\% |
| 9 | rs7848215 | 6,213,468 | C/T | 1.16 (1.12-1.20) | $2.0 \times 10^{-16}$ | 1.16 (1.13-1.19) | $2.3 \times 10^{-27}$ | 0.01 | 98\% |
| 9 | rs2066362 | 6,219,176 | G/T | 1.16 (1.11-1.21) | $6.0 \times 10^{-13}$ | 1.16 (1.13-1.20) | $9.7 \times 10^{-21}$ | 0.02 | 95\% |
| 9 | rs17582919 | 6,233,376 | T/C | 1.13 (1.09-1.17) | $1.7 \times 10^{-11}$ | 1.13 (1.10-1.16) | $7.1 \times 10^{-17}$ | 0.05 | 91\% |
| 9 | rs17498196 | 6,237,547 | A/C | 1.13 (1.09-1.17) | $5.0 \times 10^{-12}$ | 1.13 (1.10-1.16) | $7.7 \times 10^{-17}$ | 0.08 | 97\% |
| 9 | rs10815393 | 6,240,324 | T/C | 1.12 (1.08-1.17) | $4.4 \times 10^{-09}$ | 1.12 (1.09-1.16) | $7.1 \times 10^{-14}$ | 0.04 | 80\% |
| 9 | rs10491836 | 6,331,421 | C/A | 1.11 (1.07-1.15) | $2.9 \times 10^{-09}$ | 1.11 (1.08-1.14) | $9.5 \times 10^{-15}$ | 0.01 | 97\% |
| 9 | rs16924356 | 6,331,610 | G/A | 1.11 (1.07-1.15) | $2.8 \times 10^{-09}$ | 1.11 (1.08-1.14) | $1.3 \times 10^{-14}$ | 0.01 | 97\% |
| 10 | rs2589561 | 9,046,645 | A/G | 0.91 (0.88-0.94) | $3.5 \times 10^{-09}$ | 0.91 (0.88-0.94) | $3.5 \times 10^{-09}$ | 0.82 | 95\% |
| 11 | rs7130588 | 76,270,683 | A/G | 1.09 (1.06-1.12) | $3.4 \times 10^{-11}$ | 1.09 (1.06-1.12) | $2.5 \times 10^{-11}$ | 0.46 | 98\% |
| 11 | rs2155219 | 76,299,194 | G/T | 1.11 (1.08-1.15) | $5.1 \times 10^{-13}$ | 1.11 (1.08-1.14) | $1.1 \times 10^{-17}$ | 0.08 | 95\% |
| 11 | rs7927894 | 76,301,316 | C/T | 1.10 (1.08-1.13) | $2.2 \times 10^{-14}$ | 1.10 (1.08-1.13) | $2.2 \times 10^{-14}$ | 0.56 | 95\% |
| 11 | rs7927997 | 76,301,375 | C/T | 1.10 (1.07-1.13) | $3.2 \times 10^{-13}$ | 1.10 (1.07-1.13) | $3.2 \times 10^{-13}$ | 0.53 | 92\% |
| 12 | rs167769 | 57,503,775 | C/T | 1.08 (1.05-1.11) | $3.9 \times 10^{-09}$ | 1.08 (1.05-1.11) | $4.6 \times 10^{-10}$ | 0.31 | 98\% |
| 15 | rs1351544 | 61,042,867 | G/T | 0.90 (0.87-0.93) | $9.8 \times 10^{-09}$ | 0.90 (0.87-0.93) | $4.9 \times 10^{-10}$ | 0.31 | 97\% |
| 15 | rs8025324 | 61,043,378 | G/A | 0.90 (0.87-0.93) | $2.6 \times 10^{-09}$ | 0.90 (0.87-0.93) | $2.6 \times 10^{-09}$ | 0.53 | 94\% |
| 15 | rs12900122 | 61,055,411 | C/T | 0.90 (0.87-0.93) | $2.1 \times 10^{-09}$ | 0.90 (0.87-0.93) | $7.4 \times 10^{-10}$ | 0.42 | 100\% |
| 15 | rs16943087 | 61,056,035 | A/G | 0.90 (0.87-0.93) | $4.5 \times 10^{-09}$ | 0.90 (0.87-0.93) | $1.6 \times 10^{-10}$ | 0.29 | 100\% |
| 15 | rs2279292 | 61,057,770 | T/C | 0.90 (0.87-0.93) | $2.1 \times 10^{-09}$ | 0.90 (0.87-0.93) | $2.9 \times 10^{-10}$ | 0.37 | 100\% |
| 15 | rs11633029 | 61,065,553 | T/C | 0.90 (0.87-0.93) | $5.7 \times 10^{-09}$ | 0.90 (0.87-0.93) | $3.7 \times 10^{-11}$ | 0.21 | 100\% |
| 15 | rs11637671 | 61,065,607 | A/G | 0.90 (0.87-0.93) | $6.3 \times 10^{-09}$ | 0.90 (0.87-0.93) | $4.0 \times 10^{-11}$ | 0.21 | 100\% |
| 15 | rs10519067 | 61,068,347 | G/A | 0.89 (0.86-0.92) | $7.8 \times 10^{-10}$ | 0.89 (0.86-0.92) | $1.4 \times 10^{-12}$ | 0.19 | 100\% |
| 15 | rs11071558 | 61,069,421 | A/G | 0.89 (0.86-0.92) | $1.3 \times 10^{-09}$ | 0.89 (0.86-0.92) | $3.1 \times 10^{-12}$ | 0.19 | 100\% |
| 15 | rs11071559 | 61,069,988 | C/T | 0.90 (0.86-0.93) | $2.2 \times 10^{-09}$ | 0.89 (0.87-0.92) | $2.6 \times 10^{-11}$ | 0.24 | 100\% |
| 15 | rs1866316 | 67,441,997 | T/C | 1.11 (1.08-1.14) | $1.7 \times 10^{-14}$ | 1.11 (1.08-1.14) | $1.7 \times 10^{-14}$ | 0.81 | 94\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ (95 \% \mathrm{Cl})^{a} \end{array} \end{aligned}$ | $\mathbf{P r a n d o m}^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 15 | rs17293632 | 67,442,596 | C/T | 1.12 (1.08-1.15) | $1.5 \times 10^{-11}$ | 1.12 (1.09-1.15) | $1.9 \times 10^{-15}$ | 0.18 | 97\% |
| 15 | rs10152544 | 67,444,747 | C/T | 0.92 (0.90-0.94) | $1.9 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.9 \times 10^{-13}$ | 0.96 | 97\% |
| 15 | rs744910 | 67,446,785 | G/A | 0.92 (0.90-0.94) | $9.9 \times 10^{-14}$ | 0.92 (0.90-0.94) | $9.9 \times 10^{-14}$ | 0.98 | 100\% |
| 15 | rs11634793 | 67,447,452 | C/T | 0.92 (0.90-0.94) | $1.7 \times 10^{-13}$ | 0.92 (0.90-0.94) | $1.7 \times 10^{-13}$ | 0.98 | 100\% |
| 15 | rs8032739 | 67,448,899 | A/G | 1.10 (1.08-1.13) | $1.1 \times 10^{-14}$ | 1.10 (1.08-1.13) | $1.1 \times 10^{-14}$ | 0.77 | 97\% |
| 15 | rs2033784 | 67,449,660 | A/G | 1.10 (1.08-1.13) | $7.4 \times 10^{-15}$ | 1.10 (1.08-1.13) | $7.4 \times 10^{-15}$ | 0.76 | 100\% |
| 15 | rs17228058 | 67,450,305 | A/G | 1.12 (1.08-1.15) | $9.4 \times 10^{-11}$ | 1.12 (1.09-1.15) | $5.4 \times 10^{-15}$ | 0.14 | 95\% |
| 15 | rs7173698 | 67,450,893 | A/G | 1.10 (1.08-1.13) | $1.1 \times 10^{-14}$ | 1.10 (1.08-1.13) | $1.1 \times 10^{-14}$ | 0.76 | 97\% |
| 15 | rs4562997 | 67,458,152 | G/A | 1.10 (1.07-1.13) | $5.2 \times 10^{-14}$ | 1.10 (1.07-1.13) | $5.2 \times 10^{-14}$ | 0.75 | 100\% |
| 15 | rs16950687 | 67,464,013 | A/G | 1.09 (1.07-1.12) | $1.3 \times 10^{-12}$ | 1.09 (1.07-1.12) | $1.3 \times 10^{-12}$ | 0.81 | 100\% |
| 15 | rs12708492 | 67,467,541 | T/C | 0.92 (0.90-0.94) | $6.5 \times 10^{-13}$ | 0.92 (0.90-0.94) | $6.5 \times 10^{-13}$ | 0.90 | 100\% |
| 15 | rs17294280 | 67,468,285 | A/G | 1.12 (1.08-1.16) | $2.3 \times 10^{-10}$ | 1.13 (1.09-1.16) | $1.6 \times 10^{-14}$ | 0.15 | 92\% |
| 16 | rs17802927 | 11,037,738 | A/G | 0.91 (0.88-0.94) | $3.6 \times 10^{-10}$ | 0.91 (0.88-0.94) | $3.6 \times 10^{-10}$ | 0.55 | 98\% |
| 16 | rs3743976 | 11,038,824 | C/T | 0.91 (0.88-0.93) | $3.1 \times 10^{-10}$ | 0.90 (0.88-0.93) | $8.6 \times 10^{-11}$ | 0.44 | 94\% |
| 16 | rs17229044 | 11,062,936 | C/T | 0.91 (0.88-0.94) | $3.9 \times 10^{-09}$ | 0.91 (0.88-0.94) | $2.1 \times 10^{-10}$ | 0.36 | 100\% |
| 16 | rs12921922 | 11,064,321 | T/C | 0.91 (0.88-0.94) | $5.9 \times 10^{-09}$ | 0.91 (0.88-0.93) | $8.9 \times 10^{-11}$ | 0.30 | 94\% |
| 16 | rs8062923 | 11,160,966 | A/C | 1.10 (1.06-1.13) | $2.0 \times 10^{-09}$ | 1.10 (1.07-1.13) | $2.8 \times 10^{-11}$ | 0.31 | 100\% |
| 16 | rs4781035 | 11,161,178 | A/G | 1.10 (1.07-1.13) | $2.5 \times 10^{-09}$ | 1.10 (1.07-1.13) | $2.5 \times 10^{-10}$ | 0.38 | 100\% |
| 16 | rs12444495 | 11,170,455 | T/C | 1.10 (1.07-1.14) | $4.8 \times 10^{-10}$ | 1.10 (1.07-1.14) | $4.8 \times 10^{-10}$ | 0.53 | 83\% |
| 16 | rs17806299 | 11,199,980 | G/A | 0.91 (0.88-0.94) | $2.7 \times 10^{-10}$ | 0.91 (0.88-0.94) | $2.7 \times 10^{-10}$ | 0.49 | 100\% |
| 16 | rs12935657 | 11,219,041 | G/A | 0.91 (0.88-0.94) | $3.0 \times 10^{-09}$ | 0.90 (0.88-0.93) | $1.8 \times 10^{-12}$ | 0.23 | 98\% |
| 17 | rs12950186 | 37,393,395 | A/C | 0.90 (0.87-0.93) | $3.7 \times 10^{-11}$ | 0.90 (0.87-0.93) | $3.7 \times 10^{-11}$ | 0.94 | 98\% |
| 17 | rs2879258 | 37,399,379 | G/T | 0.90 (0.88-0.93) | $8.3 \times 10^{-10}$ | 0.90 (0.88-0.93) | $8.3 \times 10^{-10}$ | 0.96 | 100\% |
| 17 | rs11078895 | 37,401,051 | A/G | 0.90 (0.88-0.93) | $7.1 \times 10^{-11}$ | 0.90 (0.88-0.93) | $7.1 \times 10^{-11}$ | 0.96 | 98\% |
| 17 | rs11655972 | 37,407,072 | C/T | 0.90 (0.87-0.93) | $2.6 \times 10^{-11}$ | 0.90 (0.87-0.93) | $2.6 \times 10^{-11}$ | 0.96 | 100\% |
| 17 | rs3744349 | 37,414,842 | C/T | 0.90 (0.87-0.93) | $2.3 \times 10^{-11}$ | 0.90 (0.87-0.93) | $2.3 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs8073907 | 37,424,149 | C/T | 0.90 (0.87-0.93) | $6.7 \times 10^{-10}$ | 0.90 (0.87-0.93) | $6.7 \times 10^{-10}$ | 0.95 | 83\% |
| 17 | rs667239 | 37,442,241 | A/G | 1.11 (1.07-1.14) | $3.6 \times 10^{-11}$ | 1.11 (1.07-1.14) | $3.6 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs590051 | 37,446,571 | T/C | 1.11 (1.07-1.14) | $4.1 \times 10^{-11}$ | 1.11 (1.07-1.14) | $4.1 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs2302073 | 37,457,342 | A/G | 1.11 (1.07-1.14) | $3.7 \times 10^{-11}$ | 1.11 (1.07-1.14) | $3.7 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs584377 | 37,460,128 | G/A | 1.11 (1.07-1.14) | $3.9 \times 10^{-11}$ | 1.11 (1.07-1.14) | $3.9 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs649180 | 37,464,959 | A/C | 1.11 (1.07-1.14) | $8.3 \times 10^{-11}$ | 1.11 (1.07-1.14) | $8.3 \times 10^{-11}$ | 0.98 | 100\% |
| 17 | rs2338799 | 37,513,941 | G/A | 1.11 (1.07-1.14) | $4.0 \times 10^{-11}$ | 1.11 (1.07-1.14) | $4.0 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs6503504 | 37,514,412 | A/G | 1.11 (1.07-1.14) | $6.5 \times 10^{-11}$ | 1.11 (1.07-1.14) | $6.5 \times 10^{-11}$ | 0.97 | 97\% |
| 17 | rs9908131 | 37,520,449 | T/C | 1.11 (1.07-1.14) | $4.4 \times 10^{-11}$ | 1.11 (1.07-1.14) | $4.4 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs11078898 | 37,536,480 | A/G | 1.09 (1.06-1.12) | $8.6 \times 10^{-09}$ | 1.09 (1.06-1.11) | $4.9 \times 10^{-10}$ | 0.27 | 100\% |
| 17 | rs7208487 | 37,543,449 | T/G | 1.11 (1.07-1.14) | $6.4 \times 10^{-11}$ | 1.11 (1.07-1.14) | $6.4 \times 10^{-11}$ | 0.97 | 100\% |
| 17 | rs9906612 | 37,547,631 | A/C | 1.10 (1.07-1.14) | $6.9 \times 10^{-10}$ | 1.10 (1.07-1.14) | $6.9 \times 10^{-10}$ | 0.99 | 97\% |
| 17 | rs6503513 | 37,561,613 | A/G | 1.09 (1.06-1.12) | $5.9 \times 10^{-09}$ | 1.09 (1.06-1.12) | $5.9 \times 10^{-09}$ | 0.92 | 97\% |
| 17 | rs4795358 | 37,573,065 | A/C | 0.91 (0.88-0.93) | $2.3 \times 10^{-10}$ | 0.91 (0.88-0.93) | $2.3 \times 10^{-10}$ | 0.98 | 97\% |
| 17 | rs9646419 | 37,597,185 | A/G | 0.91 (0.88-0.93) | $2.0 \times 10^{-10}$ | 0.91 (0.88-0.93) | $2.0 \times 10^{-10}$ | 0.99 | 97\% |
| 17 | rs12938099 | 37,612,910 | T/G | 0.90 (0.88-0.93) | $1.2 \times 10^{-10}$ | 0.90 (0.88-0.93) | $1.2 \times 10^{-10}$ | 0.98 | 97\% |
| 17 | rs12449852 | 37,632,088 | A/G | 0.90 (0.87-0.93) | $5.2 \times 10^{-09}$ | 0.90 (0.87-0.93) | $5.2 \times 10^{-09}$ | 0.97 | 94\% |
| 17 | rs12936996 | 37,665,554 | G/A | 0.92 (0.89-0.95) | $8.9 \times 10^{-09}$ | 0.92 (0.90-0.94) | $4.0 \times 10^{-10}$ | 0.25 | 98\% |
| 17 | rs12937013 | 37,665,571 | G/A | 0.90 (0.87-0.93) | $6.9 \times 10^{-12}$ | 0.90 (0.87-0.93) | $6.9 \times 10^{-12}$ | 0.98 | 100\% |
| 17 | rs7503705 | 37,669,704 | A/G | 0.92 (0.89-0.94) | $2.1 \times 10^{-09}$ | 0.92 (0.89-0.94) | $1.2 \times 10^{-10}$ | 0.28 | 100\% |
| 17 | rs11078913 | 37,671,714 | G/T | 0.92 (0.89-0.94) | $8.9 \times 10^{-09}$ | 0.92 (0.89-0.94) | $2.8 \times 10^{-10}$ | 0.23 | 98\% |
| 17 | rs8065963 | 37,681,332 | C/T | 0.92 (0.89-0.94) | $3.1 \times 10^{-09}$ | 0.92 (0.89-0.94) | $1.4 \times 10^{-10}$ | 0.25 | 100\% |
| 17 | rs8069074 | 37,685,401 | A/G | 0.90 (0.87-0.93) | $4.5 \times 10^{-11}$ | 0.90 (0.87-0.93) | $4.5 \times 10^{-11}$ | 0.97 | 98\% |
| 17 | rs12450559 | 37,694,709 | G/A | 0.92 (0.89-0.94) | $2.6 \times 10^{-09}$ | 0.92 (0.89-0.94) | $1.4 \times 10^{-10}$ | 0.27 | 100\% |
| 17 | rs4239222 | 37,696,235 | G/T | 0.92 (0.89-0.95) | $3.3 \times 10^{-09}$ | 0.92 (0.90-0.94) | $2.0 \times 10^{-10}$ | 0.28 | 100\% |
| 17 | rs11657058 | 37,699,378 | T/G | 0.90 (0.87-0.93) | $8.6 \times 10^{-12}$ | 0.90 (0.87-0.93) | $8.6 \times 10^{-12}$ | 0.98 | 100\% |
| 17 | rs11657153 | 37,699,729 | A/G | 0.90 (0.87-0.93) | $2.3 \times 10^{-10}$ | 0.90 (0.87-0.93) | $2.3 \times 10^{-10}$ | 0.95 | 86\% |
| 17 | rs11654018 | 37,703,740 | T/C | 0.92 (0.89-0.94) | $2.7 \times 10^{-09}$ | 0.92 (0.90-0.94) | $2.0 \times 10^{-10}$ | 0.29 | 100\% |
| 17 | rs2303316 | 37,704,217 | A/G | 0.92 (0.89-0.95) | $9.0 \times 10^{-09}$ | 0.92 (0.90-0.94) | $4.0 \times 10^{-10}$ | 0.24 | 100\% |
| 17 | rs12947506 | 37,707,592 | T/C | 0.90 (0.87-0.93) | $1.0 \times 10^{-11}$ | 0.90 (0.87-0.93) | $1.0 \times 10^{-11}$ | 0.98 | 100\% |
| 17 | rs7503377 | 37,708,841 | T/C | 0.90 (0.87-0.93) | $2.3 \times 10^{-11}$ | 0.90 (0.87-0.93) | $2.3 \times 10^{-11}$ | 0.99 | 100\% |
| 17 | rs7216086 | 37,709,422 | G/A | 0.92 (0.89-0.95) | $6.7 \times 10^{-09}$ | 0.92 (0.90-0.94) | $3.0 \times 10^{-10}$ | 0.25 | 100\% |
| 17 | rs11078915 | 37,715,426 | T/C | 0.92 (0.89-0.94) | $8.7 \times 10^{-10}$ | 0.92 (0.89-0.94) | $7.6 \times 10^{-11}$ | 0.31 | 100\% |
| 17 | rs6503521 | 37,715,551 | G/T | 0.89 (0.87-0.92) | $1.4 \times 10^{-11}$ | 0.89 (0.87-0.92) | $1.4 \times 10^{-11}$ | 0.98 | 98\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ \text { ( } 95 \% \mathrm{Cl})^{a} \end{array} \end{aligned}$ | $\mathbf{P r a n d o m}^{\text {b }}$ | $\begin{gathered} \text { OR }_{\text {fixed }} \\ (95 \% \mathrm{Cl})^{a} \end{gathered}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathrm{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 17 | rs903507 | 37,726,423 | C/T | 1.12 (1.08-1.15) | $1.1 \times 10^{-12}$ | 1.12 (1.08-1.15) | $1.1 \times 10^{-12}$ | 0.97 | 98\% |
| 17 | rs8182252 | 37,727,950 | T/C | 1.11 (1.07-1.14) | $1.4 \times 10^{-09}$ | 1.11 (1.07-1.14) | $1.4 \times 10^{-09}$ | 0.84 | 82\% |
| 17 | rs1874226 | 37,729,031 | C/T | 1.12 (1.08-1.15) | $1.5 \times 10^{-12}$ | 1.12 (1.08-1.15) | $1.5 \times 10^{-12}$ | 0.96 | 98\% |
| 17 | rs4795385 | 37,733,148 | A/G | 1.13 (1.09-1.16) | $3.0 \times 10^{-14}$ | 1.13 (1.09-1.16) | $3.0 \times 10^{-14}$ | 0.97 | 97\% |
| 17 | rs1619021 | 37,739,274 | G/A | 0.91 (0.89-0.94) | $7.6 \times 10^{-11}$ | 0.91 (0.89-0.94) | $1.1 \times 10^{-11}$ | 0.35 | 98\% |
| 17 | rs1877030 | 37,740,161 | T/C | 0.89 (0.86-0.92) | $2.6 \times 10^{-13}$ | 0.89 (0.86-0.92) | $2.6 \times 10^{-13}$ | 0.99 | 98\% |
| 17 | rs12453198 | 37,741,879 | T/C | 0.89 (0.86-0.92) | $1.9 \times 10^{-14}$ | 0.89 (0.86-0.92) | $1.9 \times 10^{-14}$ | 0.95 | 98\% |
| 17 | rs11654954 | 37,745,979 | A/G | 0.89 (0.86-0.91) | $8.9 \times 10^{-15}$ | 0.89 (0.86-0.91) | $8.9 \times 10^{-15}$ | 0.93 | 98\% |
| 17 | rs12453682 | 37,770,005 | C/T | 0.90 (0.88-0.92) | $6.9 \times 10^{-14}$ | 0.90 (0.88-0.92) | $3.7 \times 10^{-16}$ | 0.27 | 98\% |
| 17 | rs1874228 | 37,775,274 | A/G | 0.90 (0.88-0.93) | $4.2 \times 10^{-14}$ | 0.90 (0.88-0.93) | $4.2 \times 10^{-14}$ | 0.58 | 97\% |
| 17 | rs879606 | 37,781,849 | A/G | 0.89 (0.86-0.92) | $1.4 \times 10^{-12}$ | 0.89 (0.86-0.92) | $1.4 \times 10^{-12}$ | 0.96 | 88\% |
| 17 | rs2271309 | 37,784,990 | G/A | 0.89 (0.86-0.91) | $1.9 \times 10^{-16}$ | 0.89 (0.86-0.91) | $1.9 \times 10^{-16}$ | 0.93 | 98\% |
| 17 | rs907094 | 37,790,371 | G/A | 0.89 (0.87-0.92) | $6.4 \times 10^{-15}$ | 0.89 (0.87-0.92) | $6.4 \times 10^{-15}$ | 0.91 | 98\% |
| 17 | rs3764352 | 37,790,939 | C/T | 0.90 (0.87-0.93) | $5.5 \times 10^{-11}$ | 0.90 (0.87-0.93) | $5.5 \times 10^{-11}$ | 0.91 | 80\% |
| 17 | rs9972882 | 37,807,698 | A/C | 0.88 (0.86-0.91) | $2.2 \times 10^{-21}$ | 0.88 (0.86-0.91) | $2.2 \times 10^{-21}$ | 0.78 | 98\% |
| 17 | rs1877031 | 37,814,080 | G/A | 0.89 (0.87-0.91) | $1.0 \times 10^{-21}$ | 0.89 (0.87-0.91) | $1.0 \times 10^{-21}$ | 0.61 | 98\% |
| 17 | rs2271308 | 37,817,482 | T/C | 0.89 (0.87-0.92) | $3.4 \times 10^{-18}$ | 0.89 (0.87-0.92) | $3.4 \times 10^{-18}$ | 0.81 | 98\% |
| 17 | rs931992 | 37,821,435 | G/T | 0.89 (0.87-0.91) | $2.3 \times 10^{-21}$ | 0.89 (0.87-0.91) | $2.3 \times 10^{-21}$ | 0.62 | 98\% |
| 17 | rs1053651 | 37,822,311 | A/C | 0.89 (0.87-0.92) | $1.6 \times 10^{-15}$ | 0.89 (0.87-0.92) | $1.6 \times 10^{-15}$ | 0.66 | 85\% |
| 17 | rs876493 | 37,824,545 | G/A | 0.88 (0.85-0.91) | $5.9 \times 10^{-13}$ | 0.88 (0.85-0.90) | $2.4 \times 10^{-17}$ | 0.17 | 76\% |
| 17 | rs14050 | 37,828,072 | C/T | 0.87 (0.84-0.89) | $6.3 \times 10^{-23}$ | 0.87 (0.84-0.89) | $6.3 \times 10^{-23}$ | 0.58 | 80\% |
| 17 | rs2952151 | 37,828,496 | T/C | 0.87 (0.85-0.89) | $3.8 \times 10^{-28}$ | 0.87 (0.85-0.89) | $3.8 \times 10^{-28}$ | 0.67 | 98\% |
| 17 | rs2941503 | 37,828,745 | A/G | 0.87 (0.85-0.89) | $6.8 \times 10^{-23}$ | 0.87 (0.85-0.89) | $6.8 \times 10^{-23}$ | 0.48 | 82\% |
| 17 | rs907087 | 37,828,787 | G/A | 0.87 (0.84-0.89) | $6.0 \times 10^{-23}$ | 0.87 (0.84-0.89) | $6.0 \times 10^{-23}$ | 0.51 | 80\% |
| 17 | rs903502 | 37,829,604 | C/T | 0.88 (0.86-0.90) | $1.0 \times 10^{-26}$ | 0.88 (0.86-0.90) | $1.0 \times 10^{-26}$ | 0.79 | 100\% |
| 17 | rs2941504 | 37,830,900 | A/G | 0.87 (0.85-0.89) | $1.3 \times 10^{-27}$ | 0.87 (0.85-0.89) | $1.3 \times 10^{-27}$ | 0.57 | 98\% |
| 17 | rs1565922 | 37,831,035 | A/G | 0.87 (0.85-0.89) | $7.0 \times 10^{-28}$ | 0.87 (0.85-0.89) | $7.0 \times 10^{-28}$ | 0.72 | 98\% |
| 17 | rs2934952 | 37,832,366 | G/A | 0.87 (0.85-0.89) | $1.2 \times 10^{-23}$ | 0.87 (0.85-0.89) | $1.2 \times 10^{-23}$ | 0.48 | 82\% |
| 17 | rs2941505 | 37,832,704 | A/G | 0.88 (0.86-0.90) | $2.0 \times 10^{-25}$ | 0.88 (0.86-0.90) | $2.0 \times 10^{-25}$ | 0.62 | 100\% |
| 17 | rs2941506 | 37,833,035 | A/G | 0.88 (0.86-0.90) | $1.4 \times 10^{-24}$ | 0.88 (0.86-0.90) | $1.4 \times 10^{-24}$ | 0.54 | 97\% |
| 17 | rs907089 | 37,833,600 | G/A | 0.88 (0.86-0.90) | $8.8 \times 10^{-24}$ | 0.88 (0.86-0.90) | $8.8 \times 10^{-24}$ | 0.62 | 98\% |
| 17 | rs2313171 | 37,833,842 | T/C | 0.88 (0.86-0.90) | $1.6 \times 10^{-24}$ | 0.88 (0.86-0.90) | $1.6 \times 10^{-24}$ | 0.54 | 97\% |
| 17 | rs12150298 | 37,834,541 | T/C | 0.88 (0.86-0.90) | $2.3 \times 10^{-24}$ | 0.88 (0.86-0.90) | $2.3 \times 10^{-24}$ | 0.58 | 100\% |
| 17 | rs8078228 | 37,834,998 | C/T | 0.88 (0.86-0.90) | $1.4 \times 10^{-24}$ | 0.88 (0.86-0.90) | $1.4 \times 10^{-24}$ | 0.52 | 97\% |
| 17 | rs11078919 | 37,835,755 | T/C | 0.88 (0.86-0.91) | $1.4 \times 10^{-22}$ | 0.88 (0.86-0.91) | $1.4 \times 10^{-22}$ | 0.52 | 95\% |
| 17 | rs1476278 | 37,836,243 | C/T | 0.88 (0.86-0.90) | $2.8 \times 10^{-26}$ | 0.88 (0.86-0.90) | $2.8 \times 10^{-26}$ | 0.62 | 100\% |
| 17 | rs9303274 | 37,836,353 | T/C | 0.88 (0.86-0.90) | $1.9 \times 10^{-25}$ | 0.88 (0.86-0.90) | $1.9 \times 10^{-25}$ | 0.56 | 97\% |
| 17 | rs2517957 | 37,838,716 | G/A | 0.87 (0.85-0.90) | $2.3 \times 10^{-27}$ | 0.87 (0.85-0.90) | $2.3 \times 10^{-27}$ | 0.59 | 100\% |
| 17 | rs2517958 | 37,838,751 | G/A | 0.87 (0.85-0.90) | $2.2 \times 10^{-27}$ | 0.87 (0.85-0.90) | $2.2 \times 10^{-27}$ | 0.58 | 100\% |
| 17 | rs903501 | 37,839,493 | T/C | 0.88 (0.85-0.90) | $2.0 \times 10^{-17}$ | 0.87 (0.85-0.90) | $1.1 \times 10^{-20}$ | 0.31 | 80\% |
| 17 | rs2517954 | 37,843,550 | T/C | 0.87 (0.85-0.89) | $2.1 \times 10^{-28}$ | 0.87 (0.85-0.89) | $2.1 \times 10^{-28}$ | 0.60 | 98\% |
| 17 | rs2517955 | 37,843,681 | C/T | 0.87 (0.85-0.90) | $1.3 \times 10^{-26}$ | 0.87 (0.85-0.90) | $1.3 \times 10^{-26}$ | 0.64 | 98\% |
| 17 | rs2517956 | 37,843,859 | G/A | 0.87 (0.85-0.90) | $3.3 \times 10^{-27}$ | 0.87 (0.85-0.90) | $3.3 \times 10^{-27}$ | 0.62 | 98\% |
| 17 | rs1565923 | 37,858,678 | A/G | 0.86 (0.84-0.89) | $2.1 \times 10^{-25}$ | 0.86 (0.84-0.89) | $2.1 \times 10^{-25}$ | 0.50 | 79\% |
| 17 | rs2952155 | 37,861,718 | T/C | 0.87 (0.84-0.89) | $2.9 \times 10^{-26}$ | 0.87 (0.84-0.89) | $2.9 \times 10^{-26}$ | 0.84 | 98\% |
| 17 | rs1810132 | 37,866,005 | C/T | 0.87 (0.84-0.89) | $2.9 \times 10^{-30}$ | 0.87 (0.84-0.89) | $2.9 \times 10^{-30}$ | 0.59 | 98\% |
| 17 | rs2952156 | 37,876,835 | A/G | 0.87 (0.84-0.89) | $2.2 \times 10^{-30}$ | 0.87 (0.84-0.89) | $2.2 \times 10^{-30}$ | 0.52 | 98\% |
| 17 | rs907091 | 37,921,742 | C/T | 1.18 (1.14-1.22) | $2.8 \times 10^{-20}$ | 1.18 (1.15-1.21) | $3.1 \times 10^{-36}$ | 0.01 | 86\% |
| 17 | rs907092 | 37,922,259 | G/A | 0.85 (0.83-0.88) | $2.5 \times 10^{-23}$ | 0.85 (0.83-0.87) | $3.0 \times 10^{-37}$ | 0.02 | 97\% |
| 17 | rs10445308 | 37,938,047 | C/T | 0.85 (0.83-0.88) | $1.8 \times 10^{-24}$ | 0.85 (0.83-0.87) | $7.3 \times 10^{-39}$ | 0.03 | 98\% |
| 17 | rs9909593 | 37,970,149 | A/G | 0.85 (0.83-0.88) | $5.1 \times 10^{-24}$ | 0.85 (0.83-0.87) | $7.2 \times 10^{-39}$ | 0.02 | 98\% |
| 17 | rs9303277 | 37,976,469 | C/T | 0.85 (0.83-0.88) | $9.6 \times 10^{-26}$ | 0.85 (0.83-0.87) | $4.9 \times 10^{-43}$ | 0.02 | 98\% |
| 17 | rs3816470 | 37,985,801 | A/G | 0.85 (0.83-0.88) | $1.2 \times 10^{-26}$ | 0.85 (0.83-0.87) | $6.5 \times 10^{-41}$ | 0.05 | 97\% |
| 17 | rs9635726 | 38,020,141 | $\mathrm{C} / \mathrm{T}$ | 1.16 (1.13-1.20) | $1.5 \times 10^{-23}$ | 1.16 (1.13-1.20) | $1.5 \times 10^{-23}$ | 0.52 | 95\% |
| 17 | rs4795397 | 38,023,745 | A/G | 0.86 (0.83-0.89) | $2.5 \times 10^{-16}$ | 0.86 (0.84-0.88) | $8.0 \times 10^{-29}$ | 0.01 | 82\% |
| 17 | rs11557466 | 38,024,626 | C/T | 0.85 (0.83-0.88) | $8.1 \times 10^{-22}$ | 0.85 (0.83-0.87) | $4.5 \times 10^{-36}$ | 0.02 | 95\% |
| 17 | rs11078925 | 38,025,208 | T/C | 0.85 (0.83-0.88) | $2.1 \times 10^{-24}$ | 0.85 (0.83-0.87) | $2.8 \times 10^{-38}$ | 0.03 | 98\% |
| 17 | rs12150079 | 38,025,417 | G/A | 0.88 (0.85-0.90) | $2.4 \times 10^{-15}$ | 0.88 (0.85-0.90) | $1.1 \times 10^{-21}$ | 0.08 | 97\% |
| 17 | rs11557467 | 38,028,634 | G/T | 0.85 (0.82-0.88) | $3.3 \times 10^{-21}$ | 0.85 (0.83-0.87) | $3.5 \times 10^{-31}$ | 0.07 | 80\% |
| 17 | rs10852936 | 38,031,714 | C/T | 0.86 (0.83-0.89) | $3.1 \times 10^{-17}$ | 0.86 (0.83-0.88) | $2.1 \times 10^{-27}$ | 0.03 | 80\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ (95 \% \mathrm{Cl})^{a} \end{array} \end{aligned}$ | Prandom ${ }^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathrm{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 17 | rs1054609 | 38,033,277 | A/C | 0.86 (0.83-0.88) | $1.1 \times 10^{-24}$ | 0.85 (0.83-0.88) | $3.3 \times 10^{-37}$ | 0.05 | 97\% |
| 17 | rs9907088 | 38,035,116 | G/A | 0.86 (0.83-0.88) | $1.6 \times 10^{-24}$ | 0.85 (0.83-0.87) | $6.5 \times 10^{-38}$ | 0.04 | 98\% |
| 17 | rs12232497 | 38,040,119 | T/C | 0.85 (0.82-0.88) | $1.6 \times 10^{-24}$ | 0.85 (0.83-0.87) | $9.7 \times 10^{-39}$ | 0.03 | 98\% |
| 17 | rs2872507 | 38,040,763 | G/A | 0.86 (0.83-0.88) | $2.0 \times 10^{-24}$ | 0.85 (0.83-0.87) | $8.1 \times 10^{-38}$ | 0.04 | 98\% |
| 17 | rs9901146 | 38,043,343 | G/A | 0.85 (0.82-0.87) | $9.0 \times 10^{-30}$ | 0.84 (0.83-0.87) | $6.1 \times 10^{-44}$ | 0.06 | 98\% |
| 17 | rs12950743 | 38,049,233 | T/C | 0.85 (0.82-0.87) | $6.8 \times 10^{-30}$ | 0.84 (0.82-0.87) | $5.6 \times 10^{-44}$ | 0.06 | 98\% |
| 17 | rs7359623 | 38,049,589 | C/T | 0.85 (0.82-0.87) | $1.9 \times 10^{-27}$ | 0.84 (0.82-0.87) | $2.0 \times 10^{-41}$ | 0.05 | 95\% |
| 17 | rs8067378 | 38,051,348 | A/G | 0.85 (0.83-0.88) | $1.5 \times 10^{-28}$ | 0.85 (0.83-0.87) | $4.8 \times 10^{-42}$ | 0.07 | 98\% |
| 17 | rs8069176 | 38,057,197 | G/A | 0.85 (0.82-0.88) | $2.0 \times 10^{-23}$ | 0.85 (0.83-0.87) | $4.6 \times 10^{-41}$ | 0.01 | 98\% |
| 17 | rs2305480 | 38,062,196 | G/A | 0.85 (0.82-0.87) | $3.5 \times 10^{-23}$ | 0.85 (0.83-0.87) | $2.7 \times 10^{-41}$ | 0.00 | 98\% |
| 17 | rs2305479 | 38,062,217 | C/T | 0.84 (0.81-0.87) | $2.0 \times 10^{-28}$ | 0.84 (0.82-0.86) | $1.0 \times 10^{-46}$ | 0.02 | 98\% |
| 17 | rs11078926 | 38,062,976 | G/A | 0.85 (0.81-0.88) | $2.6 \times 10^{-16}$ | 0.85 (0.82-0.87) | $5.3 \times 10^{-31}$ | 0.00 | 80\% |
| 17 | rs11078927 | 38,064,405 | C/T | 0.85 (0.82-0.87) | $1.4 \times 10^{-23}$ | 0.85 (0.82-0.87) | $2.1 \times 10^{-41}$ | 0.01 | 98\% |
| 17 | rs2290400 | 38,066,240 | T/C | 0.85 (0.82-0.88) | $1.5 \times 10^{-25}$ | 0.85 (0.83-0.87) | $1.5 \times 10^{-43}$ | 0.01 | 98\% |
| 17 | rs1008723 | 38,066,267 | G/T | 0.84 (0.82-0.87) | $6.1 \times 10^{-29}$ | 0.84 (0.82-0.86) | $9.0 \times 10^{-46}$ | 0.03 | 98\% |
| 17 | rs4795400 | 38,067,020 | C/T | 0.85 (0.82-0.88) | $7.5 \times 10^{-23}$ | 0.85 (0.83-0.87) | $5.5 \times 10^{-41}$ | 0.00 | 98\% |
| 17 | rs869402 | 38,068,043 | T/C | 1.19 (1.15-1.23) | $1.3 \times 10^{-25}$ | 1.20 (1.17-1.22) | $2.5 \times 10^{-46}$ | 0.00 | 97\% |
| 17 | rs7216389 | 38,069,949 | C/T | 1.19 (1.15-1.23) | $6.7 \times 10^{-26}$ | 1.19 (1.17-1.22) | $8.3 \times 10^{-47}$ | 0.01 | 98\% |
| 17 | rs9303280 | 38,074,031 | T/C | 1.18 (1.15-1.22) | $9.3 \times 10^{-25}$ | 1.19 (1.16-1.22) | $3.0 \times 10^{-43}$ | 0.01 | 97\% |
| 17 | rs9303281 | 38,074,046 | G/A | 1.18 (1.14-1.22) | $2.4 \times 10^{-24}$ | 1.19 (1.16-1.22) | $6.2 \times 10^{-45}$ | 0.00 | 98\% |
| 17 | rs7219923 | 38,074,518 | C/T | 1.18 (1.14-1.22) | $2.2 \times 10^{-24}$ | 1.19 (1.16-1.21) | $2.5 \times 10^{-44}$ | 0.01 | 98\% |
| 17 | rs7224129 | 38,075,426 | G/A | 1.18 (1.14-1.22) | $8.9 \times 10^{-25}$ | 1.19 (1.16-1.22) | $6.4 \times 10^{-45}$ | 0.01 | 98\% |
| 17 | rs4378650 | 38,080,865 | A/G | 1.19 (1.14-1.24) | $6.0 \times 10^{-18}$ | 1.21 (1.18-1.24) | $3.8 \times 10^{-41}$ | 0.00 | 74\% |
| 17 | rs8076131 | 38,080,912 | G/A | 1.18 (1.14-1.22) | $9.8 \times 10^{-21}$ | 1.18 (1.15-1.21) | $1.5 \times 10^{-39}$ | 0.00 | 97\% |
| 17 | rs12603332 | 38,082,807 | T/C | 1.17 (1.14-1.21) | $8.9 \times 10^{-23}$ | 1.18 (1.15-1.21) | $3.9 \times 10^{-41}$ | 0.01 | 97\% |
| 17 | rs4795402 | 38,085,385 | A/C | 1.11 (1.07-1.14) | $2.5 \times 10^{-10}$ | 1.11 (1.08-1.14) | $3.0 \times 10^{-15}$ | 0.06 | 100\% |
| 17 | rs4795405 | 38,088,417 | T/C | 1.17 (1.13-1.21) | $4.0 \times 10^{-21}$ | 1.18 (1.15-1.21) | $5.7 \times 10^{-44}$ | 0.00 | 98\% |
| 17 | rs4794820 | 38,089,344 | A/G | 1.16 (1.12-1.20) | $3.1 \times 10^{-18}$ | 1.17 (1.14-1.20) | $3.9 \times 10^{-34}$ | 0.01 | 95\% |
| 17 | rs7207600 | 38,091,660 | G/A | 1.14 (1.11-1.17) | $5.6 \times 10^{-18}$ | 1.15 (1.12-1.18) | $3.3 \times 10^{-28}$ | 0.09 | 98\% |
| 17 | rs8079416 | 38,092,713 | T/C | 1.14 (1.11-1.18) | $1.3 \times 10^{-19}$ | 1.15 (1.13-1.18) | $1.3 \times 10^{-33}$ | 0.03 | 98\% |
| 17 | rs8065126 | 38,099,035 | T/C | 1.14 (1.11-1.17) | $1.2 \times 10^{-18}$ | 1.14 (1.12-1.17) | $1.2 \times 10^{-26}$ | 0.17 | 98\% |
| 17 | rs4795408 | 38,107,627 | G/A | 1.15 (1.12-1.19) | $1.6 \times 10^{-20}$ | 1.16 (1.13-1.19) | $1.5 \times 10^{-35}$ | 0.03 | 98\% |
| 17 | rs9895948 | 38,108,363 | T/C | 1.13 (1.10-1.17) | $3.8 \times 10^{-13}$ | 1.15 (1.12-1.18) | $9.0 \times 10^{-24}$ | 0.05 | 83\% |
| 17 | rs17609240 | 38,110,689 | T/G | 1.13 (1.09-1.18) | $5.8 \times 10^{-12}$ | 1.15 (1.12-1.18) | $2.3 \times 10^{-23}$ | 0.03 | 83\% |
| 17 | rs1007654 | 38,111,354 | A/G | 1.13 (1.09-1.16) | $1.3 \times 10^{-15}$ | 1.14 (1.11-1.16) | $4.4 \times 10^{-24}$ | 0.11 | 98\% |
| 17 | rs1007655 | 38,111,419 | G/A | 1.14 (1.10-1.17) | $3.9 \times 10^{-13}$ | 1.15 (1.12-1.18) | $1.3 \times 10^{-22}$ | 0.10 | 80\% |
| 17 | rs2313640 | 38,111,845 | C/T | 1.13 (1.09-1.16) | $1.2 \times 10^{-15}$ | 1.14 (1.11-1.16) | $3.5 \times 10^{-24}$ | 0.11 | 98\% |
| 17 | rs7218742 | 38,114,361 | A/G | 1.13 (1.10-1.16) | $1.2 \times 10^{-16}$ | 1.14 (1.11-1.17) | $1.2 \times 10^{-24}$ | 0.15 | 98\% |
| 17 | rs7218321 | 38,114,469 | C/T | 1.12 (1.08-1.15) | $4.4 \times 10^{-12}$ | 1.13 (1.10-1.16) | $4.8 \times 10^{-22}$ | 0.04 | 95\% |
| 17 | rs7219080 | 38,114,516 | A/C | 1.12 (1.09-1.15) | $3.6 \times 10^{-13}$ | 1.13 (1.10-1.16) | $6.2 \times 10^{-23}$ | 0.05 | 98\% |
| 17 | rs6503526 | 38,114,598 | C/T | 1.15 (1.12-1.18) | $2.2 \times 10^{-20}$ | 1.16 (1.13-1.19) | $6.0 \times 10^{-35}$ | 0.03 | 98\% |
| 17 | rs6503527 | 38,114,719 | G/A | 1.13 (1.09-1.17) | $1.6 \times 10^{-12}$ | 1.14 (1.11-1.17) | $4.4 \times 10^{-22}$ | 0.06 | 79\% |
| 17 | rs3902025 | 38,119,254 | G/T | 1.13 (1.09-1.16) | $8.4 \times 10^{-15}$ | 1.14 (1.12-1.17) | $8.8 \times 10^{-29}$ | 0.01 | 98\% |
| 17 | rs3894194 | 38,121,993 | G/A | 1.15 (1.12-1.18) | $2.4 \times 10^{-20}$ | 1.16 (1.13-1.18) | $1.4 \times 10^{-34}$ | 0.03 | 98\% |
| 17 | rs7212938 | 38,122,680 | G/T | 0.87 (0.84-0.89) | $2.6 \times 10^{-19}$ | 0.86 (0.84-0.88) | $5.0 \times 10^{-32}$ | 0.04 | 95\% |
| 17 | rs3859192 | 38,128,648 | C/T | 1.13 (1.09-1.16) | $3.5 \times 10^{-14}$ | 1.13 (1.11-1.16) | $9.4 \times 10^{-26}$ | 0.01 | 97\% |
| 17 | rs8075668 | 38,137,623 | C/T | 0.90 (0.88-0.92) | $5.8 \times 10^{-15}$ | 0.90 (0.88-0.92) | $1.6 \times 10^{-19}$ | 0.21 | 97\% |
| 17 | rs2305481 | 38,138,624 | G/A | 0.90 (0.87-0.92) | $1.8 \times 10^{-15}$ | 0.89 (0.87-0.92) | $1.0 \times 10^{-19}$ | 0.24 | 100\% |
| 17 | rs2305482 | 38,140,927 | A/C | 0.93 (0.90-0.95) | $4.1 \times 10^{-10}$ | 0.93 (0.90-0.95) | $5.2 \times 10^{-11}$ | 0.38 | 100\% |
| 17 | rs11078930 | 38,141,955 | C/T | 0.88 (0.86-0.90) | $6.6 \times 10^{-21}$ | 0.88 (0.86-0.90) | $6.6 \times 10^{-21}$ | 0.65 | 94\% |
| 17 | rs4065321 | 38,143,548 | C/T | 0.92 (0.90-0.95) | $1.5 \times 10^{-10}$ | 0.92 (0.90-0.94) | $1.8 \times 10^{-11}$ | 0.38 | 97\% |
| 17 | rs8066582 | 38,146,929 | T/C | 0.93 (0.90-0.95) | $1.0 \times 10^{-09}$ | 0.93 (0.91-0.95) | $6.7 \times 10^{-11}$ | 0.34 | 100\% |
| 17 | rs11658328 | 38,149,236 | T/C | 0.93 (0.90-0.95) | $3.6 \times 10^{-10}$ | 0.93 (0.91-0.95) | $7.3 \times 10^{-11}$ | 0.40 | 100\% |
| 17 | rs2241245 | 38,151,014 | C/T | 0.93 (0.90-0.95) | $5.4 \times 10^{-10}$ | 0.93 (0.90-0.95) | $4.2 \times 10^{-11}$ | 0.35 | 100\% |
| 17 | rs12453334 | 38,153,473 | C/T | 0.89 (0.87-0.92) | $2.4 \times 10^{-19}$ | 0.89 (0.87-0.92) | $2.4 \times 10^{-19}$ | 0.57 | 100\% |
| 17 | rs4794822 | 38,156,712 | C/T | 1.10 (1.07-1.13) | $1.9 \times 10^{-11}$ | 1.10 (1.08-1.13) | $2.0 \times 10^{-16}$ | 0.09 | 100\% |
| 17 | rs8070454 | 38,160,754 | C/T | 1.10 (1.07-1.13) | $1.2 \times 10^{-11}$ | 1.10 (1.08-1.13) | $9.3 \times 10^{-17}$ | 0.09 | 100\% |
| 17 | rs8078723 | 38,166,879 | T/C | 1.10 (1.07-1.13) | $1.6 \times 10^{-11}$ | 1.10 (1.08-1.13) | $9.1 \times 10^{-17}$ | 0.08 | 100\% |
| 17 | rs2227319 | 38,170,845 | G/A | 0.89 (0.87-0.92) | $9.7 \times 10^{-20}$ | 0.89 (0.87-0.92) | $9.7 \times 10^{-20}$ | 0.53 | 100\% |
| 17 | rs25645 | 38,173,143 | G/A | 0.89 (0.87-0.91) | $2.1 \times 10^{-20}$ | 0.89 (0.87-0.91) | $2.1 \times 10^{-20}$ | 0.56 | 100\% |


| Chrom | SNP | Position | Allele (R/E) | $\begin{aligned} & \text { OR } \begin{array}{l} \text { random } \\ \text { ( } 95 \% \mathrm{Cl})^{\text {a }} \end{array} \end{aligned}$ | Prandom ${ }^{\text {b }}$ | $\begin{aligned} & \text { OR }_{\text {fixed }} \\ & (95 \% \mathrm{Cl})^{a} \end{aligned}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {c }}$ | $\mathrm{P}_{\text {het }}{ }^{\text {d }}$ | \% of studies contributing to metaanalysis (per SNP) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 17 | rs1042658 | 38,173,902 | C/T | 1.10 (1.07-1.13) | $3.8 \times 10^{-11}$ | 1.10 (1.08-1.13) | $1.3 \times 10^{-16}$ | 0.07 | 100\% |
| 17 | rs1045929 | 38,175,426 | C/T | 1.10 (1.07-1.14) | $3.2 \times 10^{-09}$ | 1.11 (1.08-1.14) | $1.4 \times 10^{-13}$ | 0.12 | 82\% |
| 17 | rs12309 | 38,175,462 | C/T | 0.88 (0.86-0.91) | $2.0 \times 10^{-18}$ | 0.88 (0.86-0.91) | $2.0 \times 10^{-18}$ | 0.54 | 82\% |
| 17 | rs709592 | 38,175,553 | C/T | 1.10 (1.07-1.13) | $4.4 \times 10^{-11}$ | 1.10 (1.08-1.13) | $1.4 \times 10^{-16}$ | 0.07 | 100\% |
| 17 | rs2302776 | 38,178,149 | A/G | 0.91 (0.89-0.94) | $4.5 \times 10^{-13}$ | 0.91 (0.89-0.94) | $4.5 \times 10^{-13}$ | 0.71 | 97\% |
| 17 | rs3213762 | 38,178,627 | A/G | 1.10 (1.07-1.14) | $2.2 \times 10^{-11}$ | 1.11 (1.08-1.13) | $4.8 \times 10^{-17}$ | 0.07 | 97\% |
| 17 | rs12451897 | 38,179,275 | G/T | 0.89 (0.87-0.91) | $7.9 \times 10^{-18}$ | 0.89 (0.87-0.91) | $7.9 \times 10^{-18}$ | 0.57 | 83\% |
| 17 | rs2302777 | 38,179,492 | A/G | 0.89 (0.87-0.91) | $1.1 \times 10^{-20}$ | 0.89 (0.87-0.91) | $1.1 \times 10^{-20}$ | 0.63 | 100\% |
| 17 | rs9916158 | 38,182,229 | G/T | 0.90 (0.87-0.92) | $1.1 \times 10^{-14}$ | 0.89 (0.87-0.92) | $2.1 \times 10^{-17}$ | 0.34 | 86\% |
| 17 | rs2302774 | 38,183,090 | G/T | 0.89 (0.87-0.91) | $2.8 \times 10^{-20}$ | 0.89 (0.87-0.91) | $2.8 \times 10^{-20}$ | 0.57 | 98\% |
| 17 | rs7502514 | 38,188,844 | A/G | 0.91 (0.88-0.93) | $1.5 \times 10^{-10}$ | 0.90 (0.88-0.92) | $1.9 \times 10^{-16}$ | 0.05 | 97\% |
| 17 | rs11078936 | 38,197,914 | T/C | 0.90 (0.87-0.92) | $3.5 \times 10^{-18}$ | 0.90 (0.87-0.92) | $3.5 \times 10^{-18}$ | 0.75 | 98\% |
| 17 | rs8065443 | 38,208,940 | A/G | 0.91 (0.89-0.94) | $1.7 \times 10^{-10}$ | 0.91 (0.89-0.93) | $5.5 \times 10^{-15}$ | 0.08 | 98\% |
| 17 | rs868150 | 38,213,359 | A/G | 0.91 (0.89-0.94) | $9.4 \times 10^{-11}$ | 0.91 (0.89-0.93) | $4.2 \times 10^{-15}$ | 0.10 | 98\% |
| 17 | rs17637472 | 47,461,433 | G/A | 1.08 (1.05-1.11) | $6.6 \times 10^{-09}$ | 1.08 (1.05-1.11) | $1.1 \times 10^{-09}$ | 0.35 | 98\% |

${ }^{\text {a }}$ Odds-ratios (ORs) and 95\% Confidence Intervals (CI) were computed for the effect allele (random \& fixed-effects models)
${ }^{\mathrm{b}} \mathrm{P}_{\text {random }}$ is P -value for test of association between SNP and asthma under a random-effects model
${ }^{{ }^{\text {P }} \text { fixed }}$ is $P$-value for test of association between SNP and asthma under a fixed-effects model
${ }^{d} P_{\text {het }}$ is the $P$-value for test of heterogeneity across studies with the use of Cochran's test

## Supplementary Table 9. Genetic loci associated with pediatric asthma

Results are shown for the lead SNP at each locus reaching $P_{\text {random }}<5 \times 10^{-8}$

| Region ${ }^{\text {a }}$ | Lead SNP | Position | Nearby Genes ${ }^{\text {b }}$ | Alleles $(R / E)^{c}$ | $\begin{gathered} \text { OR } \\ (95 \% \mathrm{CI})^{\mathrm{d}} \end{gathered}$ | Prandom ${ }^{\text {e }}$ | Pfixed ${ }^{\text {f }}$ | Phet ${ }^{\text {g }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2q12.1 | rs4988958 | 102,968,285 | IL1RL1,IL1RL2, IL18R1 | T/C | $\begin{gathered} 0.85 \\ (0.82-0.89) \end{gathered}$ | $5.36 \times 10^{-13}$ | $5.36 \times 10^{-13}$ | 0.78 |
| 5q31.1 | rs1295685 | 131,996,445 | IL13,RAD50,IL4 | A/G | $\begin{gathered} 0.86 \\ (0.82-0.9) \end{gathered}$ | $1.67 \times 10^{-9}$ | $1.67 \times 10^{-9}$ | 0.89 |
| 6p21.33 | rs2596464 | 31,412,961 | MICA,MICB | T/C | $\begin{gathered} 1.12 \\ (1.08-1.17) \end{gathered}$ | $1.35 \times 10^{-8}$ | $1.35 \times 10^{-8}$ | 0.47 |
| 9p24.1 | rs12551256 | 6,231,239 | IL33, RANBP6, TPD52L3 | A/G | $\begin{gathered} 0.89 \\ (0.86-0.93) \end{gathered}$ | $2.77 \times 10^{-8}$ | $2.77 \times 10^{-8}$ | 0.51 |
| 17q12-q21 | rs8069176 | 38,057,197 | ZPBP2,GSDMB | G/A | $\begin{gathered} 0.8 \\ (0.77-0.83) \\ \hline \end{gathered}$ | $4.42 \times 10^{-26}$ | $4.42 \times 10^{-26}$ | 0.77 |

[^1]Supplementary Table 10. Association of 17q12-21 SNPs with asthma in multi-ancestry and pediatric meta-analyses

|  |  |  | Multi-ancestry meta-analysis |  |  |  | Pediatric subgroup meta-analysis |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| SNP | Position | Allele (R/E) ${ }^{\text {a }}$ | OR (95\% CI) ${ }^{\text {b }}$ | Prandom ${ }^{\text {c }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ | OR ( $95 \% \mathrm{Cl})^{\text {b }}$ | $\mathbf{P r a n d o m}^{\text {c }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P}_{\text {het }}{ }^{\text {e }}$ |
| rs12950186 | 37,393,395 | A/C | 0.9 (0.87-0.93) | 3.67E-11 | 3.67E-11 | 0.94 | 0.9 (0.85-0.95) | $1.93 \mathrm{E}-04$ | 1.93E-04 | 0.72 |
| rs11078893 | 37,396,679 | T/C | 0.92 (0.89-0.95) | 8.69E-08 | 1.29E-09 | 0.20 | 0.9 (0.86-0.95) | 5.65E-05 | 5.65E-05 | 0.53 |
| rs2879258 | 37,399,379 | G/T | 0.9 (0.88-0.93) | 8.33E-10 | 8.34E-10 | 0.96 | 0.91 (0.86-0.95) | $2.55 \mathrm{E}-04$ | 2.55E-04 | 0.78 |
| rs11078895 | 37,401,051 | A/G | 0.9 (0.88-0.93) | 7.11E-11 | 7.11E-11 | 0.96 | 0.9 (0.85-0.95) | $1.42 \mathrm{E}-04$ | 1.42E-04 | 0.79 |
| rs3964723 | 37,403,979 | C/T | 0.92 (0.9-0.95) | 1.94E-08 | 7.61E-10 | 0.24 | 0.9 (0.86-0.95) | $1.76 \mathrm{E}-05$ | 1.77E-05 | 0.58 |
| rs2061342 | 37,405,657 | T/G | 0.92 (0.89-0.95) | 7.12E-08 | 1.34E-09 | 0.17 | 0.9 (0.86-0.95) | $1.39 \mathrm{E}-05$ | 1.39E-05 | 0.60 |
| rs11655972 | 37,407,072 | C/T | 0.9 (0.87-0.93) | $2.65 \mathrm{E}-11$ | 2.65E-11 | 0.96 | 0.9 (0.86-0.95) | 1.91E-04 | 1.91E-04 | 0.80 |
| rs2168785 | 37,407,135 | C/T | 0.92 (0.89-0.95) | 9.45E-08 | 1.04E-09 | 0.14 | 0.9 (0.86-0.95) | $1.73 \mathrm{E}-05$ | 1.73E-05 | 0.59 |
| rs755500 | 37,409,865 | G/A | 0.92 (0.9-0.95) | $1.55 \mathrm{E}-08$ | $4.91 \mathrm{E}-10$ | 0.23 | 0.9 (0.86-0.95) | $1.38 \mathrm{E}-05$ | 1.38E-05 | 0.60 |
| rs752314 | 37,410,700 | C/T | 0.92 (0.89-0.95) | $1.42 \mathrm{E}-08$ | 3.59E-10 | 0.21 | 0.9 (0.86-0.95) | $1.42 \mathrm{E}-05$ | 1.42E-05 | 0.59 |
| rs3744349 | 37,414,842 | C/T | 0.9 (0.88-0.93) | 2.29E-11 | 2.29E-11 | 0.97 | 0.9 (0.85-0.95) | 9.87E-05 | 9.87E-05 | 0.82 |
| rs11078897 | 37,415,870 | C/T | 0.92 (0.89-0.95) | 1.59E-08 | 4.02E-10 | 0.21 | 0.9 (0.86-0.95) | $1.48 \mathrm{E}-05$ | 1.48E-05 | 0.59 |
| rs2338755 | 37,419,317 | T/C | 0.92 (0.89-0.95) | $1.68 \mathrm{E}-08$ | 4.11E-10 | 0.21 | 0.9 (0.86-0.95) | $1.51 \mathrm{E}-05$ | 1.51E-05 | 0.59 |
| rs8073907 | 37,424,149 | C/T | 0.9 (0.87-0.93) | 6.73E-10 | 6.73E-10 | 0.95 | 0.9 (0.85-0.95) | $4.08 \mathrm{E}-04$ | 4.08E-04 | 0.82 |
| rs8079590 | 37,426,201 | T/C | 0.91 (0.88-0.95) | 6.72E-07 | 4.85E-09 | 0.12 | 0.9 (0.85-0.95) | $1.04 \mathrm{E}-04$ | 4.64E-05 | 0.39 |
| rs4795355 | 37,427,382 | C/T | 0.92 (0.89-0.95) | $4.51 \mathrm{E}-08$ | 8.17E-10 | 0.18 | 0.9 (0.86-0.95) | $1.45 \mathrm{E}-05$ | 1.45E-05 | 0.60 |
| rs600010 | 37,433,517 | т/C | 0.92 (0.89-0.95) | 5.11E-08 | $8.45 \mathrm{E}-10$ | 0.17 | 0.9 (0.86-0.95) | $1.48 \mathrm{E}-05$ | 1.48E-05 | 0.60 |
| rs602688 | 37,439,496 | т/C | 1.09 (1.05-1.12) | 5.02E-08 | 8.09E-10 | 0.17 | 1.11 (1.06-1.16) | $1.57 \mathrm{E}-05$ | 1.57E-05 | 0.60 |
| rs588193 | 37,440,439 | A/C | 1.09 (1.06-1.12) | 2.07E-08 | $4.65 \mathrm{E}-10$ | 0.20 | 1.11 (1.06-1.16) | $1.70 \mathrm{E}-05$ | 1.70E-05 | 0.59 |
| rs801426 | 37,441,109 | G/A | 1.09 (1.06-1.12) | 2.67E-08 | 4.42E-10 | 0.18 | 1.11 (1.06-1.16) | $1.69 \mathrm{E}-05$ | 1.69E-05 | 0.60 |
| rs667239 | 37,442,241 | A/G | 1.11 (1.08-1.14) | 3.57E-11 | 3.57E-11 | 0.97 | 1.11 (1.05-1.17) | $1.94 \mathrm{E}-04$ | 1.94E-04 | 0.80 |
| rs590051 | 37,446,571 | T/C | 1.11 (1.07-1.14) | 4.06E-11 | 4.06E-11 | 0.97 | 1.11 (1.05-1.17) | $1.73 \mathrm{E}-04$ | 1.73E-04 | 0.83 |
| rs6322 | 37,453,294 | C/A | 1.09 (1.06-1.12) | 7.42E-08 | 1.19E-09 | 0.16 | 1.11 (1.06-1.16) | $1.27 \mathrm{E}-05$ | 1.27E-05 | 0.61 |
| rs620686 | 37,453,617 | A/G | 1.09 (1.06-1.12) | 2.08E-08 | 4.86E-10 | 0.20 | 1.11 (1.06-1.16) | $1.61 \mathrm{E}-05$ | 1.61E-05 | 0.60 |
| rs2302073 | 37,457,342 | A/G | 1.11 (1.08-1.14) | $3.74 \mathrm{E}-11$ | $3.74 \mathrm{E}-11$ | 0.97 | 1.11 (1.05-1.17) | $1.79 \mathrm{E}-04$ | 1.79E-04 | 0.83 |
| rs584377 | 37,460,128 | G/A | 1.11 (1.08-1.14) | 3.87E-11 | 3.87E-11 | 0.97 | 1.11 (1.05-1.17) | $1.79 \mathrm{E}-04$ | 1.79E-04 | 0.84 |
| rs10491129 | 37,461,643 | T/G | 1.09 (1.05-1.12) | 6.00E-08 | 9.20E-10 | 0.16 | 1.11 (1.06-1.16) | $1.48 \mathrm{E}-05$ | 1.48E-05 | 0.61 |
| rs649180 | 37,464,959 | A/C | 1.11 (1.07-1.14) | 8.28E-11 | 8.28E-11 | 0.98 | 1.11 (1.05-1.17) | $1.86 \mathrm{E}-04$ | 1.86E-04 | 0.83 |
| rs9904334 | 37,468,336 | G/A | 1.09 (1.06-1.12) | 2.07E-08 | 5.25E-10 | 0.21 | 1.11 (1.06-1.16) | $1.64 \mathrm{E}-05$ | 1.64E-05 | 0.60 |
| rs8066704 | 37,477,218 | C/T | 1.09 (1.06-1.12) | $1.83 \mathrm{E}-08$ | 4.19E-10 | 0.20 | 1.11 (1.06-1.16) | $1.68 \mathrm{E}-05$ | 1.68E-05 | 0.60 |
| rs8070695 | 37,478,062 | G/A | 1.09 (1.05-1.12) | 5.45E-08 | 9.86E-10 | 0.17 | 1.11 (1.06-1.16) | $1.53 \mathrm{E}-05$ | 1.53E-05 | 0.60 |
| rs9944411 | 37,483,599 | T/C | 1.09 (1.05-1.12) | $2.78 \mathrm{E}-08$ | 7.32E-10 | 0.21 | 1.11 (1.06-1.16) | $1.57 \mathrm{E}-05$ | 1.58E-05 | 0.60 |
| rs7220650 | 37,487,168 | т/C | 1.09 (1.06-1.12) | 1.41E-08 | $3.45 \mathrm{E}-10$ | 0.21 | 1.11 (1.06-1.16) | $1.62 \mathrm{E}-05$ | 1.62E-05 | 0.60 |
| rs9916302 | 37,499,949 | т/C | 1.09 (1.05-1.12) | 2.83E-08 | 8.44E-10 | 0.22 | 1.11 (1.06-1.16) | $1.64 \mathrm{E}-05$ | 1.64E-05 | 0.60 |
| rs8069451 | 37,504,933 | T/C | 1.09 (1.06-1.12) | 1.97E-08 | 6.38E-10 | 0.23 | 1.11 (1.06-1.17) | 5.45E-06 | 5.45E-06 | 0.57 |
| rs9894586 | 37,505,421 | C/T | 1.09 (1.06-1.12) | 3.79E-08 | 5.24E-10 | 0.19 | 1.11 (1.06-1.16) | $2.04 \mathrm{E}-05$ | 2.04E-05 | 0.55 |
| rs9892055 | 37,510,402 | C/T | 1.09 (1.06-1.12) | 2.63E-08 | 7.66E-10 | 0.21 | 1.11 (1.06-1.16) | $1.52 \mathrm{E}-05$ | 1.52E-05 | 0.61 |
| rs2338799 | 37,513,941 | G/A | 1.11 (1.07-1.14) | 4.03E-11 | $4.03 \mathrm{E}-11$ | 0.97 | 1.11 (1.05-1.17) | $1.96 \mathrm{E}-04$ | 1.96E-04 | 0.84 |
| rs6503503 | 37,514,373 | G/A | 1.09 (1.05-1.12) | $2.13 \mathrm{E}-08$ | 8.52E-10 | 0.24 | 1.11 (1.06-1.16) | $1.66 \mathrm{E}-05$ | 1.66E-05 | 0.60 |
| rs6503504 | 37,514,412 | A/G | 1.11 (1.07-1.14) | 6.48E-11 | 6.48E-11 | 0.97 | 1.11 (1.05-1.17) | 2.02E-04 | $2.02 \mathrm{E}-04$ | 0.84 |
| rs8076494 | 37,516,722 | T/C | 1.09 (1.05-1.12) | 1.99E-08 | 7.94E-10 | 0.24 | 1.11 (1.06-1.16) | $1.59 \mathrm{E}-05$ | 1.59E-05 | 0.60 |
| rs9908131 | 37,520,449 | T/C | 1.11 (1.07-1.14) | 4.41E-11 | $4.41 \mathrm{E}-11$ | 0.97 | 1.11 (1.05-1.17) | $1.96 \mathrm{E}-04$ | 1.96E-04 | 0.84 |
| rs7223438 | 37,521,355 | т/C | 1.09 (1.05-1.12) | 6.32E-08 | 1.46E-09 | 0.19 | 1.11 (1.06-1.16) | $1.65 \mathrm{E}-05$ | 1.65E-05 | 0.60 |
| rs6503507 | 37,525,274 | C/T | 1.09 (1.06-1.12) | $2.55 \mathrm{E}-08$ | 8.31E-10 | 0.22 | 1.11 (1.06-1.16) | $1.61 \mathrm{E}-05$ | 1.61E-05 | 0.60 |
| rs11078898 | 37,536,480 | A/G | 1.09 (1.06-1.12) | 8.58E-09 | $4.88 \mathrm{E}-10$ | 0.27 | 1.11 (1.06-1.16) | $1.59 \mathrm{E}-05$ | 1.59E-05 | 0.61 |
| rs7221875 | 37,543,328 | G/A | 1.09 (1.05-1.12) | 6.63E-08 | 2.11E-09 | 0.20 | 1.11 (1.06-1.16) | $1.18 \mathrm{E}-05$ | 1.18E-05 | 0.61 |
| rs7208487 | 37,543,449 | T/G | 1.11 (1.07-1.14) | 6.42E-11 | 6.42E-11 | 0.97 | 1.1 (1.05-1.16) | 2.92E-04 | 2.92E-04 | 0.82 |
| rs9906612 | 37,547,631 | A/C | 1.1 (1.07-1.14) | 6.92E-10 | 6.92E-10 | 0.99 | 1.11 (1.05-1.17) | $1.98 \mathrm{E}-04$ | 1.98E-04 | 0.84 |
| rs10491128 | 37,551,988 | G/A | 1.08 (1.05-1.11) | 5.94E-08 | 3.06E-09 | 0.25 | 1.11 (1.06-1.16) | $1.42 \mathrm{E}-05$ | 1.42E-05 | 0.60 |
| rs6503513 | 37,561,613 | A/G | 1.09 (1.06-1.12) | 5.85E-09 | 5.85E-09 | 0.92 | 1.11 (1.05-1.17) | 7.08E-05 | 7.08E-05 | 0.95 |
| rs4795357 | 37,569,551 | T/C | 0.92 (0.9-0.95) | 5.98E-08 | $2.74 \mathrm{E}-09$ | 0.24 | 0.9 (0.86-0.94) | $1.08 \mathrm{E}-05$ | 1.08E-05 | 0.61 |
| rs4795358 | 37,573,065 | A/C | 0.91 (0.88-0.93) | $2.28 \mathrm{E}-10$ | $2.28 \mathrm{E}-10$ | 0.98 | 0.9 (0.86-0.95) | 1.17E-04 | 1.17E-04 | 0.85 |
| rs7501488 | 37,576,417 | T/G | 0.92 (0.9-0.95) | 2.87E-08 | $1.44 \mathrm{E}-09$ | 0.25 | 0.9 (0.86-0.94) | $1.02 \mathrm{E}-05$ | 1.02E-05 | 0.60 |
| rs10445306 | 37,591,422 | A/G | 0.92 (0.89-0.95) | 1.06E-08 | 3.77E-10 | 0.25 | 0.9 (0.86-0.94) | 7.67E-06 | 7.67E-06 | 0.60 |
| rs9646419 | 37,597,185 | A/G | 0.91 (0.88-0.93) | $1.99 \mathrm{E}-10$ | $1.99 \mathrm{E}-10$ | 0.99 | 0.9 (0.85-0.95) | $9.57 \mathrm{E}-05$ | 9.57E-05 | 0.84 |
| rs4795369 | 37,609,120 | G/A | 0.93 (0.9-0.95) | 2.76E-07 | $2.62 \mathrm{E}-08$ | 0.24 | 0.9 (0.86-0.94) | $8.53 \mathrm{E}-06$ | 8.53E-06 | 0.6 |


|  |  |  | Multi-ancestry meta-analysis |  |  |  | Pediatric subgroup meta-analysis |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| SNP | Position | Allele (R/E) ${ }^{\text {a }}$ | OR ( $95 \% \mathrm{CI})^{\text {b }}$ | Prandom ${ }^{\text {c }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathrm{Phet}^{\text {e }}$ | OR (95\% CI) ${ }^{\text {b }}$ | $\mathrm{Prandom}^{\text {c }}$ | Pfixed ${ }^{\text {d }}$ | $\mathbf{P r e t}^{\text {e }}$ |
| rs12938099 | 37,612,910 | T/G | 0.9 (0.88-0.93) | 1.18E-10 | 1.18E-10 | 0.98 | 0.9 (0.85-0.95) | 7.40E-05 | 7.40E-05 | 0.85 |
| rs4390625 | 37,620,347 | A/G | 0.92 (0.89-0.95) | 6.00E-08 | 1.50E-09 | 0.19 | 0.9 (0.86-0.94) | 7.13E-06 | 7.13E-06 | 0.57 |
| rs11078901 | 37,625,912 | C/T | 0.92 (0.89-0.95) | 5.55E-08 | 9.90E-10 | 0.21 | 0.9 (0.86-0.94) | 9.62E-06 | 9.62E-06 | 0.51 |
| rs12449852 | 37,632,088 | A/G | 0.9 (0.87-0.93) | 5.19E-09 | 5.19E-09 | 0.97 | 0.9 (0.85-0.95) | 3.64E-04 | 3.64E-04 | 0.71 |
| rs12948906 | 37,644,854 | C/T | 0.91 (0.88-0.95) | 8.71E-07 | $1.23 \mathrm{E}-08$ | 0.15 | 0.89 (0.85-0.94) | 5.91E-05 | 2.39E-05 | 0.39 |
| rs11870631 | 37,645,129 | C/T | 0.92 (0.89-0.95) | 5.01E-08 | 1.63E-09 | 0.21 | 0.9 (0.86-0.94) | 6.95E-06 | 6.95E-06 | 0.58 |
| rs2018026 | 37,659,606 | G/A | 0.92 (0.89-0.95) | 5.56E-08 | 8.89E-10 | 0.20 | 0.9 (0.86-0.94) | 9.16E-06 | 9.16E-06 | 0.53 |
| rs6503518 | 37,660,469 | A/G | 0.91 (0.88-0.95) | 8.09E-07 | 1.13E-08 | 0.15 | 0.89 (0.85-0.94) | 5.41E-05 | $2.36 \mathrm{E}-05$ | 0.40 |
| rs11078907 | 37,662,954 | C/T | 0.92 (0.89-0.95) | $3.22 \mathrm{E}-08$ | 1.11E-09 | 0.23 | 0.9 (0.86-0.94) | 6.95E-06 | 6.95E-06 | 0.59 |
| rs7225096 | 37,664,443 | T/C | 0.92 (0.89-0.95) | $3.76 \mathrm{E}-08$ | 1.29E-09 | 0.23 | 0.9 (0.86-0.94) | 7.27E-06 | 7.27E-06 | 0.59 |
| rs12936996 | 37,665,554 | G/A | 0.92 (0.89-0.95) | 8.90E-09 | 4.02E-10 | 0.25 | 0.9 (0.86-0.94) | 7.23E-06 | 7.23E-06 | 0.59 |
| rs12937013 | 37,665,571 | G/A | 0.9 (0.87-0.93) | 6.86E-12 | 6.86E-12 | 0.98 | 0.9 (0.85-0.95) | 6.69E-05 | 6.70E-05 | 0.83 |
| rs7503705 | 37,669,704 | A/G | 0.92 (0.89-0.94) | 2.15E-09 | 1.16E-10 | 0.28 | 0.9 (0.86-0.94) | 5.36E-06 | 5.36E-06 | 0.62 |
| rs11078913 | 37,671,714 | G/T | 0.92 (0.89-0.94) | 8.88E-09 | $2.83 \mathrm{E}-10$ | 0.23 | 0.9 (0.86-0.94) | 3.73E-06 | 3.73E-06 | 0.62 |
| rs11658678 | 37,680,096 | C/T | 0.91 (0.89-0.94) | 1.05E-08 | 1.74E-10 | 0.22 | 0.9 (0.86-0.94) | 4.51E-06 | 4.51E-06 | 0.57 |
| rs8065963 | 37,681,332 | C/T | 0.92 (0.89-0.94) | 3.08E-09 | 1.39E-10 | 0.25 | 0.9 (0.86-0.94) | 4.76E-06 | 4.76E-06 | 0.62 |
| rs8069074 | 37,685,401 | A/G | 0.9 (0.87-0.93) | $4.53 \mathrm{E}-11$ | 4.53E-11 | 0.97 | 0.9 (0.85-0.95) | 5.47E-05 | 5.47E-05 | 0.82 |
| rs12450559 | 37,694,709 | G/A | 0.92 (0.89-0.94) | 2.58E-09 | 1.41E-10 | 0.27 | 0.9 (0.86-0.94) | 4.57E-06 | 4.57E-06 | 0.60 |
| rs4239222 | 37,696,235 | G/T | 0.92 (0.89-0.95) | 3.29E-09 | 2.05E-10 | 0.28 | 0.9 (0.86-0.94) | 4.16E-06 | 4.16E-06 | 0.61 |
| rs11657058 | 37,699,378 | T/G | 0.9 (0.87-0.93) | 8.60E-12 | 8.60E-12 | 0.98 | 0.9 (0.85-0.95) | 6.66E-05 | 6.66E-05 | 0.80 |
| rs11657153 | 37,699,729 | A/G | 0.9 (0.87-0.93) | $2.28 \mathrm{E}-10$ | 2.28E-10 | 0.95 | 0.89 (0.84-0.95) | 6.16E-04 | 6.16E-04 | 0.66 |
| rs11654018 | 37,703,740 | T/C | 0.92 (0.89-0.94) | $2.67 \mathrm{E}-09$ | 2.04E-10 | 0.29 | 0.9 (0.86-0.94) | 5.63E-06 | 5.63E-06 | 0.58 |
| rs2303316 | 37,704,217 | A/G | 0.92 (0.89-0.95) | 9.04E-09 | 3.97E-10 | 0.24 | 0.9 (0.86-0.94) | 4.59E-06 | 4.59E-06 | 0.61 |
| rs12947506 | 37,707,592 | T/C | 0.9 (0.87-0.93) | 1.03E-11 | 1.03E-11 | 0.98 | 0.9 (0.85-0.95) | 8.79E-05 | 8.79E-05 | 0.80 |
| rs7503377 | 37,708,841 | т/C | 0.9 (0.87-0.93) | $2.27 \mathrm{E}-11$ | 2.27E-11 | 0.99 | 0.9 (0.85-0.95) | 8.57E-05 | 8.58E-05 | 0.79 |
| rs7216086 | 37,709,422 | G/A | 0.92 (0.89-0.95) | 6.74E-09 | 3.01E-10 | 0.25 | 0.9 (0.86-0.94) | 5.42E-06 | 5.42E-06 | 0.60 |
| rs11078915 | 37,715,426 | T/C | 0.92 (0.89-0.94) | 8.69E-10 | 7.62E-11 | 0.31 | 0.9 (0.86-0.94) | 2.90E-06 | 2.90E-06 | 0.60 |
| rs6503521 | 37,715,551 | G/T | 0.89 (0.87-0.92) | $1.37 \mathrm{E}-11$ | 1.37E-11 | 0.98 | 0.89 (0.85-0.94) | 5.72E-05 | 5.72E-05 | 0.76 |
| rs7503195 | 37,722,515 | G/A | 0.93 (0.91-0.96) | $2.53 \mathrm{E}-07$ | $2.53 \mathrm{E}-07$ | 0.52 | 0.93 (0.89-0.97) | 1.41E-03 | 1.41E-03 | 0.52 |
| rs903507 | 37,726,423 | C/T | 1.12 (1.09-1.15) | $1.13 \mathrm{E}-12$ | 1.13E-12 | 0.97 | 1.13 (1.07-1.19) | 2.70E-05 | $2.70 \mathrm{E}-05$ | 0.80 |
| rs8182252 | 37,727,950 | T/C | 1.11 (1.07-1.15) | 1.43E-09 | 1.43E-09 | 0.84 | 1.13 (1.07-1.2) | 4.10E-05 | 4.10E-05 | 0.92 |
| rs1874226 | 37,729,031 | C/T | 1.12 (1.08-1.15) | $1.51 \mathrm{E}-12$ | 1.51E-12 | 0.96 | 1.13 (1.07-1.19) | $2.05 \mathrm{E}-05$ | $2.05 \mathrm{E}-05$ | 0.80 |
| rs4795385 | 37,733,148 | A/G | 1.13 (1.09-1.16) | $2.95 \mathrm{E}-14$ | 2.95E-14 | 0.97 | 1.15 (1.09-1.21) | 5.91E-07 | 5.91E-07 | 0.80 |
| rs1619021 | 37,739,274 | G/A | 0.91 (0.89-0.94) | 7.61E-11 | 1.09E-11 | 0.35 | 0.89 (0.85-0.93) | 1.02E-06 | 1.02E-06 | 0.65 |
| rs1877030 | 37,740,161 | T/C | 0.89 (0.86-0.92) | $2.55 \mathrm{E}-13$ | 2.56E-13 | 0.99 | 0.88 (0.84-0.93) | $1.11 \mathrm{E}-0$ | 1.11E-05 | 0.80 |
| rs12453198 | 37,741,879 | T/C | 0.89 (0.86-0.92) | 1.93E-14 | 1.93E-14 | 0.95 | 0.89 (0.84-0.93) | 7.84E-06 | 7.84E-06 | 0.51 |
| rs11654954 | 37,745,979 | A/G | 0.89 (0.86-0.91) | 8.91E-15 | 8.91E-15 | 0.93 | 0.88 (0.84-0.93) | 5.31E-06 | 5.31E-06 | 0.53 |
| rs12453682 | 37,770,005 | C/T | 0.9 (0.88-0.92) | $6.89 \mathrm{E}-14$ | 3.75E-16 | 0.27 | 0.87 (0.83-0.91) | 7.29E-10 | $9.40 \mathrm{E}-11$ | 0.37 |
| rs1874228 | 37,775,274 | A/G | 0.9 (0.88-0.93) | 4.22E-14 | 4.23E-14 | 0.58 | 0.87 (0.83-0.91) | 4.28E-09 | 4.28E-09 | 0.67 |
| rs879606 | 37,781,849 | A/G | 0.89 (0.86-0.92) | 1.36E-12 | 1.36E-12 | 0.96 | 0.88 (0.83-0.93) | 4.29E-06 | 4.29E-06 | 0.78 |
| rs2271309 | 37,784,990 | G/A | 0.89 (0.86-0.91) | $1.95 \mathrm{E}-16$ | 1.95E-16 | 0.93 | 0.87 (0.83-0.91) | 7.98E-09 | 7.98E-09 | 0.88 |
| rs907094 | 37,790,371 | G/A | 0.89 (0.87-0.92) | 6.37E-15 | 6.37E-15 | 0.91 | 0.87 (0.83-0.91) | 8.33E-09 | 8.33E-09 | 0.85 |
| rs3764352 | 37,790,939 | C/T | 0.9 (0.87-0.93) | 5.45E-11 | 5.45E-11 | 0.91 | 0.86 (0.82-0.91) | 4.31E-07 | 4.31E-07 | 0.96 |
| rs9972882 | 37,807,698 | A/C | 0.88 (0.86-0.91) | 2.19E-21 | 2.19E-21 | 0.78 | 0.86 (0.82-0.9) | 7.40E-12 | 7.40E-12 | 0.54 |
| rs1877031 | 37,814,080 | G/A | 0.89 (0.87-0.91) | 1.05E-21 | 1.05E-21 | 0.61 | 0.85 (0.82-0.89) | 6.25E-14 | 6.26E-14 | 0.82 |
| rs2271308 | 37,817,482 | T/C | 0.89 (0.87-0.92) | $3.42 \mathrm{E}-18$ | $3.42 \mathrm{E}-18$ | 0.81 | 0.86 (0.83-0.9) | $1.28 \mathrm{E}-11$ | $1.28 \mathrm{E}-11$ | 0.79 |
| rs931992 | 37,821,435 | G/T | 0.89 (0.87-0.91) | $2.26 \mathrm{E}-21$ | 2.26E-21 | 0.62 | 0.85 (0.82-0.89) | $7.28 \mathrm{E}-14$ | 7.28E-14 | 0.84 |
| rs1053651 | 37,822,311 | A/C | 0.89 (0.87-0.92) | $1.62 \mathrm{E}-15$ | 1.62E-15 | 0.66 | 0.85 (0.8-0.89) | 1.01E-09 | 1.01E-09 | 0.78 |
| rs876493 | 37,824,545 | G/A | 0.88 (0.85-0.91) | 5.89E-13 | 2.44E-17 | 0.17 | 0.83 (0.79-0.88) | 1.05E-10 | 3.58E-11 | 0.42 |
| rs14050 | 37,828,072 | C/T | 0.87 (0.84-0.89) | 6.28E-23 | 6.29E-23 | 0.58 | 0.83 (0.79-0.88) | 6.42E-12 | $2.14 \mathrm{E}-13$ | 0.35 |
| rs2952151 | 37,828,496 | T/C | 0.87 (0.85-0.89) | $3.78 \mathrm{E}-28$ | 3.78E-28 | 0.67 | 0.84 (0.81-0.88) | 3.89E-15 | 6.94E-16 | 0.41 |
| rs2941503 | 37,828,745 | A/G | 0.87 (0.85-0.89) | 6.79E-23 | 6.79E-23 | 0.48 | 0.83 (0.79-0.88) | $3.22 \mathrm{E}-12$ | $1.77 \mathrm{E}-13$ | 0.37 |
| rs907087 | 37,828,787 | G/A | 0.87 (0.84-0.89) | 6.00E-23 | 6.00E-23 | 0.51 | 0.83 (0.79-0.88) | $4.82 \mathrm{E}-12$ | $1.65 \mathrm{E}-13$ | 0.35 |
| rs903502 | 37,829,604 | C/T | 0.88 (0.86-0.9) | $1.00 \mathrm{E}-26$ | 1.00E-26 | 0.79 | 0.84 (0.81-0.88) | 1.03E-15 | 1.03E-15 | 0.68 |
| rs2941504 | 37,830,900 | A/G | 0.87 (0.85-0.89) | 1.28E-27 | 1.28E-27 | 0.57 | 0.84 (0.81-0.88) | 1.53E-14 | 1.50E-15 | 0.39 |
| rs1565922 | 37,831,035 | A/G | 0.87 (0.85-0.89) | 6.99E-28 | 6.99E-28 | 0.72 | 0.84 (0.81-0.88) | 1.82E-14 | 1.18E-15 | 0.38 |
| rs2934952 | 37,832,366 | G/A | 0.87 (0.85-0.89) | 1.19E-23 | 1.19E-23 | 0.48 | 0.83 (0.79-0.88) | 5.56E-10 | 8.84E-13 | 0.24 |
| rs2941505 | 37,832,704 | A/G | 0.88 (0.86-0.9) | $1.98 \mathrm{E}-25$ | 1.98E-25 | 0.62 | 0.84 (0.81-0.88) | 1.06E-15 | 1.06E-15 | 0.60 |
| rs2941506 | 37,833,035 | A/G | 0.88 (0.86-0.9) | $1.42 \mathrm{E}-24$ | 1.42E-24 | 0.54 | 0.84 (0.81-0.88) | 1.11E-15 | $1.11 \mathrm{E}-15$ | 0.59 |


|  |  |  | Multi-ancestry meta-analysis |  |  |  | Pediatric subgroup meta-analysis |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| SNP | Position | Allele (R/E) ${ }^{\text {a }}$ | OR (95\% CI) ${ }^{\text {b }}$ | $\mathrm{Prandom}^{\text {c }}$ | $\mathbf{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathrm{P}_{\text {het }}{ }^{\text {e }}$ | OR (95\% CI) ${ }^{\text {b }}$ | $\mathrm{Prandom}^{\text {c }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {d }}$ | $\mathbf{P h e t}^{\text {e }}$ |
| rs907089 | 37,833,600 | G/A | 0.88 (0.86-0.9) | $8.78 \mathrm{E}-24$ | 8.78E-24 | 0.62 | 0.84 (0.81-0.88) | $1.31 \mathrm{E}-15$ | 1.31E-15 | 0.62 |
| rs2313171 | 37,833,842 | T/C | 0.88 (0.86-0.9) | $1.57 \mathrm{E}-24$ | $1.57 \mathrm{E}-24$ | 0.54 | 0.84 (0.81-0.88) | $1.33 \mathrm{E}-15$ | $1.33 \mathrm{E}-15$ | 0.59 |
| rs12150298 | 37,834,541 | T/C | 0.88 (0.86-0.9) | $2.33 \mathrm{E}-24$ | 2.33E-24 | 0.58 | 0.85 (0.81-0.88) | $2.23 \mathrm{E}-15$ | 2.23E-15 | 0.58 |
| rs8078228 | 37,834,998 | C/T | 0.88 (0.86-0.9) | $1.38 \mathrm{E}-24$ | $1.38 \mathrm{E}-24$ | 0.52 | 0.84 (0.81-0.88) | $1.11 \mathrm{E}-15$ | $1.11 \mathrm{E}-15$ | 0.57 |
| rs11078919 | 37,835,755 | T/C | 0.88 (0.86-0.91) | $1.43 \mathrm{E}-22$ | $1.43 \mathrm{E}-22$ | 0.52 | 0.84 (0.81-0.88) | $1.23 \mathrm{E}-15$ | $1.23 \mathrm{E}-15$ | 0.57 |
| rs1476278 | 37,836,243 | C/T | 0.88 (0.86-0.9) | $2.82 \mathrm{E}-26$ | 2.82E-26 | 0.62 | 0.84 (0.81-0.88) | $1.28 \mathrm{E}-15$ | $1.28 \mathrm{E}-15$ | 0.57 |
| rs9303274 | 37,836,353 | T/C | 0.88 (0.86-0.9) | 1.87E-25 | $1.87 \mathrm{E}-25$ | 0.56 | 0.84 (0.81-0.88) | $1.26 \mathrm{E}-15$ | 1.27E-15 | 0.56 |
| rs2517957 | 37,838,716 | G/A | 0.87 (0.85-0.9) | $2.31 \mathrm{E}-27$ | 2.31E-27 | 0.59 | 0.84 (0.81-0.88) | 9.79E-16 | 9.79E-16 | 0.57 |
| rs2517958 | 37,838,751 | G/A | 0.87 (0.85-0.9) | $2.18 \mathrm{E}-27$ | $2.18 \mathrm{E}-27$ | 0.58 | 0.84 (0.81-0.88) | $9.28 \mathrm{E}-16$ | 9.28E-16 | 0.57 |
| rs903501 | 37,839,493 | T/C | 0.88 (0.85-0.9) | $1.96 \mathrm{E}-17$ | $1.07 \mathrm{E}-20$ | 0.31 | 0.83 (0.78-0.88) | $1.55 \mathrm{E}-09$ | 5.29E-12 | 0.26 |
| rs2517954 | 37,843,550 | T/C | 0.87 (0.85-0.89) | $2.09 \mathrm{E}-28$ | 2.09E-28 | 0.60 | 0.84 (0.81-0.88) | $3.76 \mathrm{E}-15$ | $3.76 \mathrm{E}-15$ | 0.51 |
| rs2517955 | 37,843,681 | C/T | 0.87 (0.85-0.9) | $1.26 \mathrm{E}-26$ | $1.26 \mathrm{E}-26$ | 0.64 | 0.85 (0.81-0.88) | $7.27 \mathrm{E}-15$ | 7.27E-15 | 0.70 |
| rs2517956 | 37,843,859 | G/A | 0.87 (0.85-0.9) | $3.31 \mathrm{E}-27$ | $3.31 \mathrm{E}-27$ | 0.62 | 0.84 (0.81-0.88) | $5.77 \mathrm{E}-15$ | 5.77E-15 | 0.49 |
| rs1565923 | 37,858,678 | A/G | 0.86 (0.84-0.89) | $2.09 \mathrm{E}-25$ | $2.09 \mathrm{E}-25$ | 0.50 | 0.84 (0.79-0.89) | $5.15 \mathrm{E}-10$ | $1.28 \mathrm{E}-11$ | 0.32 |
| rs2952155 | 37,861,718 | T/C | 0.87 (0.84-0.89) | $2.92 \mathrm{E}-26$ | $2.92 \mathrm{E}-26$ | 0.84 | 0.85 (0.81-0.89) | $1.93 \mathrm{E}-12$ | $1.93 \mathrm{E}-12$ | 0.74 |
| rs1810132 | 37,866,005 | C/T | 0.87 (0.84-0.89) | $2.94 \mathrm{E}-30$ | 2.94E-30 | 0.59 | 0.84 (0.81-0.88) | $2.42 \mathrm{E}-15$ | $2.42 \mathrm{E}-15$ | 0.50 |
| rs2952156 | 37,876,835 | A/G | 0.87 (0.84-0.89) | $2.24 \mathrm{E}-30$ | $2.24 \mathrm{E}-30$ | 0.52 | 0.84 (0.81-0.88) | $2.42 \mathrm{E}-15$ | $2.42 \mathrm{E}-15$ | 0.51 |
| rs907091 | 37,921,742 | C/T | 1.18 (1.14-1.22) | $2.82 \mathrm{E}-20$ | 3.14E-36 | 0.01 | 1.26 (1.2-1.32) | $5.59 \mathrm{E}-20$ | $1.03 \mathrm{E}-21$ | 0.38 |
| rs907092 | 37,922,259 | G/A | 0.85 (0.83-0.88) | $2.52 \mathrm{E}-23$ | 2.96E-37 | 0.02 | 0.81 (0.78-0.84) | $3.73 \mathrm{E}-24$ | 3.73E-24 | 0.69 |
| rs10445308 | 37,938,047 | C/T | 0.85 (0.83-0.88) | $1.76 \mathrm{E}-24$ | 7.29E-39 | 0.03 | 0.81 (0.77-0.84) | $1.55 \mathrm{E}-24$ | $1.55 \mathrm{E}-24$ | 0.80 |
| rs9909593 | 37,970,149 | A/G | 0.85 (0.83-0.88) | $5.13 \mathrm{E}-24$ | 7.22E-39 | 0.02 | 0.81 (0.77-0.84) | $8.23 \mathrm{E}-25$ | 8.23E-25 | 0.85 |
| rs9303277 | 37,976,469 | C/T | 0.85 (0.83-0.88) | $9.57 \mathrm{E}-26$ | $4.88 \mathrm{E}-43$ | 0.02 | 0.8 (0.77-0.84) | $1.24 \mathrm{E}-25$ | $1.24 \mathrm{E}-25$ | 0.62 |
| rs3816470 | 37,985,801 | A/G | 0.85 (0.83-0.88) | $1.23 \mathrm{E}-26$ | 6.46E-41 | 0.05 | 0.81 (0.78-0.85) | 8.83E-22 | 8.83E-22 | 0.60 |
| rs9635726 | 38,020,141 | C/T | 1.16 (1.13-1.2) | $1.50 \mathrm{E}-23$ | 1.50E-23 | 0.52 | 1.17 (1.12-1.23) | $4.53 \mathrm{E}-11$ | $4.53 \mathrm{E}-11$ | 0.57 |
| rs4795397 | 38,023,745 | A/G | 0.86 (0.83-0.89) | $2.55 \mathrm{E}-16$ | 7.99E-29 | 0.01 | 0.8 (0.76-0.84) | $1.64 \mathrm{E}-19$ | $1.64 \mathrm{E}-19$ | 0.79 |
| rs11557466 | 38,024,626 | C/T | 0.85 (0.83-0.88) | 8.06E-22 | 4.55E-36 | 0.02 | 0.81 (0.77-0.84) | 6.37E-24 | 6.37E-24 | 0.83 |
| rs11078925 | 38,025,208 | T/C | 0.85 (0.83-0.88) | $2.15 \mathrm{E}-24$ | $2.82 \mathrm{E}-38$ | 0.03 | 0.81 (0.77-0.84) | $2.36 \mathrm{E}-24$ | 2.36E-24 | 0.87 |
| rs12150079 | 38,025,417 | G/A | 0.88 (0.85-0.9) | $2.45 \mathrm{E}-15$ | $1.14 \mathrm{E}-21$ | 0.08 | 0.82 (0.79-0.86) | 7.87E-17 | 7.87E-17 | 0.65 |
| rs11557467 | 38,028,634 | G/T | 0.85 (0.82-0.88) | $3.27 \mathrm{E}-21$ | 3.50E-31 | 0.07 | 0.8 (0.76-0.84) | $4.02 \mathrm{E}-21$ | 4.02E-21 | 0.51 |
| rs10852936 | 38,031,714 | C/T | 0.86 (0.83-0.89) | $3.10 \mathrm{E}-17$ | $2.13 \mathrm{E}-27$ | 0.03 | 0.8 (0.76-0.84) | 6.80E-21 | 6.80E-21 | 0.81 |
| rs1054609 | 38,033,277 | A/C | 0.86 (0.83-0.88) | $1.14 \mathrm{E}-24$ | 3.27E-37 | 0.05 | 0.8 (0.77-0.84) | 7.56E-25 | 7.56E-25 | 0.88 |
| rs9907088 | 38,035,116 | G/A | 0.86 (0.83-0.88) | $1.58 \mathrm{E}-24$ | 6.50E-38 | 0.04 | 0.81 (0.77-0.84) | $9.80 \mathrm{E}-25$ | 9.80E-25 | 0.86 |
| rs12232497 | 38,040,119 | T/C | 0.85 (0.83-0.88) | $1.57 \mathrm{E}-24$ | $9.74 \mathrm{E}-39$ | 0.03 | 0.8 (0.77-0.84) | $9.16 \mathrm{E}-25$ | 9.16E-25 | 0.86 |
| rs2872507 | 38,040,763 | G/A | 0.86 (0.83-0.88) | $2.02 \mathrm{E}-24$ | 8.06E-38 | 0.04 | 0.81 (0.77-0.84) | $1.65 \mathrm{E}-24$ | $1.65 \mathrm{E}-24$ | 0.86 |
| rs9901146 | 38,043,343 | G/A | 0.85 (0.82-0.87) | $9.03 \mathrm{E}-30$ | 6.09E-44 | 0.06 | 0.81 (0.77-0.84) | $3.37 \mathrm{E}-25$ | $3.37 \mathrm{E}-25$ | 0.63 |
| rs12950743 | 38,049,233 | T/C | 0.85 (0.82-0.87) | 6.78E-30 | $5.58 \mathrm{E}-44$ | 0.06 | 0.81 (0.77-0.84) | $3.00 \mathrm{E}-25$ | 3.00E-25 | 0.62 |
| rs7359623 | 38,049,589 | C/T | 0.85 (0.82-0.87) | $1.93 \mathrm{E}-27$ | $2.03 \mathrm{E}-41$ | 0.05 | 0.8 (0.77-0.84) | $1.46 \mathrm{E}-24$ | $1.46 \mathrm{E}-24$ | 0.67 |
| rs8067378 | 38,051,348 | A/G | 0.85 (0.83-0.88) | $1.46 \mathrm{E}-28$ | $4.78 \mathrm{E}-42$ | 0.07 | 0.81 (0.77-0.84) | 6.58E-25 | 6.58E-25 | 0.62 |
| rs8069176 | 38,057,197 | G/A | 0.85 (0.82-0.88) | $1.97 \mathrm{E}-23$ | $4.55 \mathrm{E}-41$ | 0.01 | 0.8 (0.77-0.83) | $4.42 \mathrm{E}-26$ | 4.42E-26 | 0.77 |
| rs2305480 | 38,062,196 | G/A | 0.85 (0.82-0.88) | $3.47 \mathrm{E}-23$ | $2.72 \mathrm{E}-41$ | $4.6 \times 10^{-3}$ | 0.8 (0.77-0.83) | $5.37 \mathrm{E}-26$ | 5.37E-26 | 0.74 |
| rs2305479 | 38,062,217 | C/T | 0.84 (0.81-0.87) | $2.00 \mathrm{E}-28$ | $1.03 \mathrm{E}-46$ | 0.02 | 0.8 (0.77-0.84) | 6.20E-26 | 6.21E-26 | 0.51 |
| rs11078926 | 38,062,976 | G/A | 0.85 (0.81-0.88) | $2.57 \mathrm{E}-16$ | 5.28E-31 | $1.8 \times 10^{-3}$ | 0.79 (0.75-0.83) | 7.56E-22 | 7.56E-22 | 0.75 |
| rs11078927 | 38,064,405 | C/T | 0.85 (0.82-0.87) | $1.45 \mathrm{E}-23$ | $2.14 \mathrm{E}-41$ | 0.01 | 0.8 (0.77-0.84) | $1.78 \mathrm{E}-25$ | $1.78 \mathrm{E}-25$ | 0.79 |
| rs2290400 | 38,066,240 | T/C | 0.85 (0.82-0.88) | $1.51 \mathrm{E}-25$ | $1.46 \mathrm{E}-43$ | 0.01 | 0.8 (0.77-0.84) | 7.81E-25 | 7.81E-25 | 0.51 |
| rs1008723 | 38,066,267 | G/T | 0.84 (0.82-0.87) | 6.13E-29 | 8.99E-46 | 0.03 | 0.8 (0.77-0.84) | 4.96E-26 | 4.96E-26 | 0.48 |
| rs4795400 | 38,067,020 | C/T | 0.85 (0.82-0.88) | $7.48 \mathrm{E}-23$ | $5.51 \mathrm{E}-41$ | $3.6 \times 10^{-3}$ | 0.8 (0.77-0.84) | $1.51 \mathrm{E}-25$ | 1.51E-25 | 0.76 |
| rs869402 | 38,068,043 | T/C | 1.19 (1.15-1.23) | $1.27 \mathrm{E}-25$ | $2.52 \mathrm{E}-46$ | $4.4 \times 10^{-3}$ | 1.24 (1.19-1.3) | $3.97 \mathrm{E}-24$ | 6.38E-25 | 0.43 |
| rs7216389 | 38,069,949 | C/T | 1.19 (1.15-1.23) | 6.73E-26 | 8.27E-47 | 0.01 | 1.25 (1.19-1.3) | 7.97E-23 | 7.17E-26 | 0.32 |
| rs9303280 | 38,074,031 | T/C | 1.18 (1.15-1.22) | $9.30 \mathrm{E}-25$ | 2.96E-43 | 0.01 | 1.24 (1.19-1.3) | $3.43 \mathrm{E}-21$ | 6.91E-25 | 0.28 |
| rs9303281 | 38,074,046 | G/A | 1.18 (1.14-1.22) | $2.41 \mathrm{E}-24$ | 6.20E-45 | $4.4 \times 10^{-3}$ | 1.25 (1.19-1.3) | $3.60 \mathrm{E}-23$ | 5.35E-26 | 0.33 |
| rs7219923 | 38,074,518 | C/T | 1.18 (1.14-1.22) | $2.20 \mathrm{E}-24$ | $2.47 \mathrm{E}-44$ | 0.01 | 1.25 (1.19-1.3) | $3.46 \mathrm{E}-23$ | 5.52E-26 | 0.33 |
| rs7224129 | 38,075,426 | G/A | 1.18 (1.14-1.22) | $8.94 \mathrm{E}-25$ | $6.40 \mathrm{E}-45$ | 0.01 | 1.25 (1.2-1.31) | $1.29 \mathrm{E}-24$ | 1.30E-26 | 0.38 |
| rs4378650 | 38,080,865 | A/G | 1.19 (1.14-1.24) | 5.99E-18 | $3.78 \mathrm{E}-41$ | $2.9 \times 10^{-3}$ | 1.23 (1.17-1.3) | 5.39E-15 | $1.01 \mathrm{E}-19$ | 0.18 |
| rs8076131 | 38,080,912 | G/A | 1.18 (1.14-1.22) | $9.81 \mathrm{E}-21$ | $1.48 \mathrm{E}-39$ | $1.7 \times 10^{-3}$ | 1.25 (1.2-1.3) | $6.18 \mathrm{E}-25$ | 6.18E-25 | 0.72 |
| rs12603332 | 38,082,807 | T/C | 1.17 (1.14-1.21) | 8.90E-23 | $3.87 \mathrm{E}-41$ | 0.01 | 1.23 (1.18-1.29) | $2.41 \mathrm{E}-20$ | $1.54 \mathrm{E}-23$ | 0.30 |
| rs17608925 | 38,082,831 | T/C | 0.9 (0.87-0.94) | $3.26 \mathrm{E}-07$ | 3.26E-07 | 0.48 | 0.87 (0.81-0.94) | $2.35 \mathrm{E}-04$ | 2.35E-04 | 0.85 |
| rs3744246 | 38,084,350 | T/C | 1.11 (1.07-1.15) | 5.08E-08 | 1.06E-13 | 0.01 | 1.16 (1.1-1.22) | 8.22E-09 | 8.22E-09 | 0.70 |
| rs4795402 | 38,085,385 | A/C | 1.11 (1.07-1.14) | $2.50 \mathrm{E}-10$ | $2.98 \mathrm{E}-15$ | 0.06 | 1.14 (1.09-1.2) | $3.38 \mathrm{E}-08$ | $2.56 \mathrm{E}-08$ | 0.45 |
| rs4795403 | 38,085,722 | T/C | 1.11 (1.07-1.16) | $1.50 \mathrm{E}-07$ | 2.20E-13 | $2.7 \times 10^{-3}$ | 1.16 (1.1-1.22) | $1.31 \mathrm{E}-08$ | $1.31 \mathrm{E}-08$ | 0.62 |


|  |  |  | Multi-ancestry meta-analysis |  |  |  | Pediatric subgroup meta-analysis |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| SNP | Position | Allele (R/E) ${ }^{\text {a }}$ | OR (95\% CI) ${ }^{\text {b }}$ | Prandom ${ }^{\text {c }}$ | $\mathrm{P}_{\text {fixed }}{ }^{\text {d }}$ | Phet ${ }^{\text {e }}$ | OR (95\% CI) ${ }^{\text {b }}$ | $\mathrm{Prandom}^{\text {c }}$ | $P_{\text {fixed }}{ }^{\text {d }}$ | $\mathrm{Phet}^{\text {e }}$ |
| rs4795404 | 38,085,791 | A/C | 1.11 (1.07-1.15) | 6.14E-08 | 1.40E-13 | 0.01 | 1.16 (1.1-1.22) | 9.06E-09 | 9.06E-09 | 0.68 |
| rs4795405 | 38,088,417 | T/C | 1.17 (1.13-1.21) | 4.05E-21 | 5.74E-44 | $1.4 \times 10^{-3}$ | 1.23 (1.18-1.28) | $2.33 \mathrm{E}-22$ | 2.33E-22 | 0.63 |
| rs4794820 | 38,089,344 | A/G | 1.16 (1.12-1.2) | 3.12E-18 | 3.89E-34 | 0.01 | 1.24 (1.19-1.3) | 7.18E-23 | 7.18E-23 | 0.79 |
| rs7207600 | 38,091,660 | G/A | 1.14 (1.11-1.17) | 5.63E-18 | 3.34E-28 | 0.09 | 1.19 (1.14-1.25) | $2.54 \mathrm{E}-15$ | $2.54 \mathrm{E}-15$ | 0.53 |
| rs8079416 | 38,092,713 | T/C | 1.15 (1.11-1.18) | 1.28E-19 | 1.28E-33 | 0.03 | 1.2 (1.15-1.25) | 4.84E-17 | 6.83E-18 | 0.40 |
| rs8065126 | 38,099,035 | T/C | 1.14 (1.11-1.17) | 1.19E-18 | $1.25 \mathrm{E}-26$ | 0.17 | 1.19 (1.14-1.24) | 8.54E-15 | 8.54E-15 | 0.56 |
| rs4795408 | 38,107,627 | G/A | 1.15 (1.12-1.19) | 1.59E-20 | $1.54 \mathrm{E}-35$ | 0.03 | 1.19 (1.14-1.25) | $8.71 \mathrm{E}-16$ | $2.57 \mathrm{E}-17$ | 0.34 |
| rs9895948 | 38,108,363 | T/C | 1.14 (1.1-1.17) | 3.76E-13 | 9.00E-24 | 0.05 | 1.22 (1.16-1.28) | 5.46E-14 | 5.46E-14 | 0.75 |
| rs17609240 | 38,110,689 | T/G | 1.13 (1.09-1.18) | 5.76E-12 | 2.30E-23 | 0.03 | 1.22 (1.16-1.29) | $2.92 \mathrm{E}-14$ | 2.93E-14 | 0.75 |
| rs1007654 | 38,111,354 | A/G | 1.13 (1.09-1.16) | 1.29E-15 | 4.36E-24 | 0.11 | 1.17 (1.12-1.22) | $1.94 \mathrm{E}-13$ | 1.94E-13 | 0.47 |
| rs1007655 | 38,111,419 | G/A | 1.14 (1.1-1.18) | $3.89 \mathrm{E}-13$ | 1.28E-22 | 0.10 | 1.22 (1.15-1.28) | 7.76E-14 | 7.76E-14 | 0.79 |
| rs2313640 | 38,111,845 | C/T | 1.13 (1.09-1.16) | 1.20E-15 | 3.51E-24 | 0.11 | 1.17 (1.12-1.22) | $2.27 \mathrm{E}-13$ | $2.27 \mathrm{E}-13$ | 0.47 |
| rs7218742 | 38,114,361 | A/G | 1.13 (1.1-1.16) | 1.18E-16 | 1.23E-24 | 0.15 | 1.18 (1.13-1.23) | $1.61 \mathrm{E}-13$ | 1.61E-13 | 0.50 |
| rs7218321 | 38,114,469 | C/T | 1.12 (1.08-1.15) | 4.44E-12 | $4.81 \mathrm{E}-22$ | 0.04 | 1.17 (1.13-1.23) | $1.82 \mathrm{E}-13$ | 1.82E-13 | 0.47 |
| rs7219080 | 38,114,516 | A/C | 1.12 (1.09-1.15) | $3.59 \mathrm{E}-13$ | 6.21E-23 | 0.05 | 1.17 (1.12-1.23) | 4.12E-13 | 2.08E-13 | 0.43 |
| rs6503526 | 38,114,598 | C/T | 1.15 (1.12-1.18) | 2.17E-20 | 5.96E-35 | 0.03 | 1.19 (1.14-1.24) | 3.90E-16 | 5.36E-17 | 0.40 |
| rs6503527 | 38,114,719 | G/A | 1.13 (1.09-1.17) | 1.59E-12 | 4.41E-22 | 0.06 | 1.18 (1.12-1.24) | 5.45E-11 | $4.74 \mathrm{E}-12$ | 0.34 |
| rs3902025 | 38,119,254 | G/T | 1.13 (1.1-1.16) | $8.44 \mathrm{E}-15$ | 8.79E-29 | 0.01 | 1.18 (1.13-1.23) | $1.71 \mathrm{E}-15$ | 1.71E-15 | 0.80 |
| rs3894194 | 38,121,993 | G/A | 1.15 (1.12-1.18) | $2.43 \mathrm{E}-20$ | 1.45E-34 | 0.03 | 1.2 (1.15-1.25) | $8.21 \mathrm{E}-18$ | 2.62E-18 | 0.43 |
| rs7212938 | 38,122,680 | G/T | 0.87 (0.84-0.89) | $2.56 \mathrm{E}-19$ | 4.95E-32 | 0.04 | 0.82 (0.78-0.86) | $1.70 \mathrm{E}-15$ | 1.89E-18 | 0.24 |
| rs3859192 | 38,128,648 | C/T | 1.13 (1.09-1.16) | $3.50 \mathrm{E}-14$ | 9.38E-26 | 0.01 | 1.17 (1.12-1.23) | $1.94 \mathrm{E}-10$ | 3.70E-14 | 0.09 |
| rs8075668 | 38,137,623 | C/T | 0.9 (0.88-0.92) | 5.83E-15 | 1.58E-19 | 0.21 | 0.88 (0.84-0.92) | $4.01 \mathrm{E}-08$ | 6.42E-10 | 0.20 |
| rs2305481 | 38,138,624 | G/A | 0.9 (0.87-0.92) | 1.84E-15 | 1.01E-19 | 0.24 | 0.89 (0.85-0.93) | $2.62 \mathrm{E}-07$ | 3.20E-08 | 0.29 |
| rs2305482 | 38,140,927 | A/C | 0.93 (0.9-0.95) | $4.09 \mathrm{E}-10$ | 5.23E-11 | 0.38 | 0.91 (0.88-0.95) | $1.77 \mathrm{E}-05$ | 1.77E-05 | 0.64 |
| rs11078930 | 38,141,955 | C/T | 0.88 (0.86-0.9) | 6.64E-21 | 6.64E-21 | 0.65 | 0.87 (0.83-0.91) | 1.00E-09 | 1.00E-09 | 0.47 |
| rs4065321 | 38,143,548 | C/T | 0.92 (0.9-0.95) | $1.51 \mathrm{E}-10$ | 1.75E-11 | 0.38 | 0.91 (0.88-0.95) | $1.52 \mathrm{E}-05$ | 1.52E-05 | 0.61 |
| rs8066582 | 38,146,929 | т/C | 0.93 (0.9-0.95) | 1.02E-09 | 6.68E-11 | 0.34 | 0.91 (0.88-0.95) | $1.56 \mathrm{E}-05$ | 1.56E-05 | 0.60 |
| rs11658328 | 38,149,236 | T/C | 0.93 (0.91-0.95) | $3.60 \mathrm{E}-10$ | 7.31E-11 | 0.40 | 0.92 (0.88-0.95) | $1.89 \mathrm{E}-05$ | 1.89E-05 | 0.65 |
| rs2241245 | 38,151,014 | C/T | 0.93 (0.9-0.95) | 5.38E-10 | 4.20E-11 | 0.36 | 0.91 (0.88-0.95) | 1.53E-05 | 1.53E-05 | 0.65 |
| rs12453334 | 38,153,473 | C/T | 0.89 (0.87-0.92) | $2.43 \mathrm{E}-19$ | 2.43E-19 | 0.57 | 0.88 (0.85-0.92) | 3.54E-09 | 3.19E-09 | 0.46 |
| rs4794822 | 38,156,712 | C/T | 1.1 (1.07-1.13) | 1.89E-11 | 1.96E-16 | 0.09 | 1.13 (1.08-1.17) | 1.02E-08 | 1.02E-08 | 0.57 |
| rs8070454 | 38,160,754 | C/T | 1.1 (1.07-1.13) | $1.24 \mathrm{E}-11$ | 9.29E-17 | 0.09 | 1.13 (1.08-1.17) | 9.77E-09 | 9.77E-09 | 0.58 |
| rs8078723 | 38,166,879 | T/C | 1.1 (1.07-1.13) | 1.56E-11 | 9.13E-17 | 0.08 | 1.13 (1.08-1.18) | 9.46E-09 | 9.46E-09 | 0.59 |
| rs2227319 | 38,170,845 | G/A | 0.89 (0.87-0.92) | 9.69E-20 | 9.69E-20 | 0.53 | 0.88 (0.85-0.92) | 9.38E-09 | 4.97E-09 | 0.42 |
| rs25645 | 38,173,143 | G/A | 0.89 (0.87-0.91) | 2.05E-20 | 2.05E-20 | 0.56 | 0.88 (0.84-0.92) | 3.98E-09 | 3.98E-09 | 0.55 |
| rs1042658 | 38,173,902 | C/T | 1.1 (1.07-1.13) | 3.83E-11 | 1.26E-16 | 0.07 | 1.13 (1.08-1.17) | $1.49 \mathrm{E}-08$ | 1.49E-08 | 0.57 |
| rs1045929 | 38,175,426 | C/T | 1.1 (1.07-1.14) | 3.25E-09 | 1.36E-13 | 0.12 | 1.16 (1.1-1.22) | 3.71E-09 | 3.71E-09 | 0.72 |
| rs12309 | 38,175,462 | C/T | 0.88 (0.86-0.91) | $2.02 \mathrm{E}-18$ | 2.02E-18 | 0.54 | 0.86 (0.82-0.91) | 3.16E-09 | 3.16E-09 | 0.69 |
| rs709592 | 38,175,553 | C/T | 1.1 (1.07-1.13) | 4.40E-11 | 1.45E-16 | 0.07 | 1.13 (1.08-1.17) | $1.55 \mathrm{E}-08$ | 1.55E-08 | 0.57 |
| rs2302776 | 38,178,149 | A/G | 0.91 (0.89-0.94) | $4.47 \mathrm{E}-13$ | 4.47E-13 | 0.71 | 0.91 (0.87-0.95) | 3.48E-06 | $3.48 \mathrm{E}-06$ | 0.63 |
| rs3213762 | 38,178,627 | A/G | 1.1 (1.07-1.14) | 2.20E-11 | 4.77E-17 | 0.07 | 1.13 (1.08-1.17) | 1.98E-08 | 1.98E-08 | 0.55 |
| rs12451897 | 38,179,275 | G/T | 0.89 (0.87-0.91) | 7.88E-18 | 7.88E-18 | 0.57 | 0.87 (0.82-0.91) | 1.95E-08 | 1.95E-08 | 0.62 |
| rs2302777 | 38,179,492 | A/G | 0.89 (0.87-0.91) | 1.13E-20 | 1.13E-20 | 0.63 | 0.88 (0.85-0.92) | 6.24E-09 | 6.24E-09 | 0.49 |
| rs9916158 | 38,182,229 | G/T | 0.9 (0.87-0.92) | 1.07E-14 | 2.06E-17 | 0.34 | 0.88 (0.84-0.93) | 1.85E-06 | 2.98E-07 | 0.31 |
| rs2302774 | 38,183,090 | G/T | 0.89 (0.87-0.91) | 2.79E-20 | 2.79E-20 | 0.57 | 0.89 (0.85-0.93) | $2.46 \mathrm{E}-08$ | 2.46E-08 | 0.47 |
| rs7502514 | 38,188,844 | A/G | 0.91 (0.88-0.93) | 1.50E-10 | 1.88E-16 | 0.05 | 0.89 (0.85-0.93) | $5.31 \mathrm{E}-08$ | 5.31E-08 | 0.57 |
| rs3935280 | 38,189,055 | A/G | 0.91 (0.87-0.94) | 1.05E-07 | 6.35E-13 | 0.05 | 0.86 (0.81-0.9) | $4.78 \mathrm{E}-09$ | $4.78 \mathrm{E}-09$ | 0.78 |
| rs11078936 | 38,197,914 | T/C | 0.9 (0.87-0.92) | $3.52 \mathrm{E}-18$ | 3.52E-18 | 0.75 | 0.88 (0.85-0.92) | 6.53E-09 | 6.53E-09 | 0.51 |
| rs8065443 | 38,208,940 | A/G | 0.91 (0.89-0.94) | 1.73E-10 | 5.47E-15 | 0.08 | 0.89 (0.85-0.93) | $2.45 \mathrm{E}-08$ | $2.45 \mathrm{E}-08$ | 0.46 |
| rs868150 | 38,213,359 | A/G | 0.91 (0.89-0.94) | $9.43 \mathrm{E}-11$ | $4.23 \mathrm{E}-15$ | 0.10 | 0.89 (0.85-0.93) | $2.03 \mathrm{E}-08$ | 2.03E-08 | 0.50 |
| rs7502966 | 38,216,522 | C/T | 1.07 (1.04-1.1) | 1.52E-07 | $1.52 \mathrm{E}-07$ | 0.85 | 1.08 (1.03-1.13) | 1.04E-03 | 1.04E-03 | 0.63 |

[^2]Supplementary Table 11. Association between cis-gene transcripts in blood and lung and 17q12-21 SNPs
a. Association between lead asthma SNPs (rs2952156 in multi-ancestry meta-analysis and rs8069176 in pediatric metaanalysis) and cis-gene transcripts

${ }^{a}$ The gene where eventually the lead SNP lies is first indicated followed by the previous and next genes
${ }^{\text {b }}$ eQTL databases are described in Online Methods
${ }^{\text {cBlood }}$ is whole blood; LCL = lymphoblastoid cell lines; Lung is whole lung tissue
${ }^{d} P_{\text {transcript }}$ is the $P$-value for test of association of asthma-associated SNP with gene transcript

## b. Association analysis of ORMDL3 and PGAP3 transcripts with their most significantly associated SNPs (peak SNPs) and the 17q12-21 asthma-associated SNPs

This analysis focused on ORMDL3 transcript in blood (showing the strongest association with both 17q12-21 asthma SNPs, rs2952156 and rs8069176, in the majority of five blood eQTL databases) and PGAP3 transcript in lung (associated with rs2952156), as shown above. It used the summary statistics from the GTEx eQTL-database, available in both blood ( $\mathrm{n}=338$ samples) and lung ( $\mathrm{n}=278$ samples).

We first performed association analysis of these gene transcripts with their respective peak SNPs and then by conditioning on the lead asthmaassociated SNPs (using the GCTA software (PMID 23756893 ) and the large pan-European ECRHS study as the reference to estimate LD between SNPs, similarly to the approximate conditional analysis conducted for asthma and explained in detail in the Online Methods).

As can be seen in the following table, rs8069176 accounts for a large part of the association of the peak SNP (rs9303281) with ORMDL3 transcript in blood while rs2952156 accounts for a large part of the association of the peak SNP (rs2941505) with PGAP3 transcript in lung.

| Tissue | Gene_transcript | Association of lead asthma SNP with gene transcript |  |  | Association of peak SNP with gene transcript |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | Asthma SNP | Position | $P_{\text {transcript }}{ }^{\text {a }}$ | Peak SNP ${ }^{\text {b }}$ | Position | $\mathbf{r a c}^{\mathbf{2 c}}$ | Ptranscript ${ }^{\text {d }}$ | $\mathbf{P e}_{\text {adj for }}$ asthma SNP |
| Blood | ORMDL3 | rs2952156 | 37,876,835 | $3.7 \times 10^{-6}$ | rs9303281 | 38,074,046 | 0.39 | $6.3 \times 10^{-30}$ | $7.6 \times 10^{-23}$ |
|  |  | rs8069176 | 38,057,197 | $1.2 \times 10^{-26}$ | rs9303281 | 38,074,046 | 0.79 | $6.3 \times 10^{-30}$ | $2.2 \times 10^{-4}$ |
| Lung | PGAP3 | rs2952156 | 37,876,835 | $1.0 \times 10^{-9}$ | rs2941505 | 37,832,704 | 0.84 | $5.6 \times 10^{-12}$ | $1.5 \times 10^{-3}$ |
|  |  | rs8069176 | 38,057,197 | $1.4 \times 10^{-3}$ | rs2941505 | 37,832,704 | 0.35 | $5.6 \times 10^{-12}$ | $2.6 \times 10^{-9}$ |

${ }^{2} \mathrm{P}$-value for association of the asthma SNP with gene transcript
${ }^{\mathrm{b}}$ Most significant SNP (Peak SNP) associated with the gene transcript
${ }^{\text {c Correlation }}\left(\mathrm{r}^{2}\right)$ between the asthma SNP and the peak SNP
${ }^{d P}$-value for association of the peak SNP with gene transcript
${ }^{e} P$-value for association of the peak SNP with gene transcript while conditioning on the asthma SNP

Supplementary Table 12. Asthma loci reported in the GWAS catalog wih $\mathrm{P}<5 \times 10^{-8}$ and not replicated in TAGC meta-analyses

| Information from TAGC meta-analysis Information from GWAS Catalog |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| REGION | Position build 37 | TAGC SNP | $\begin{gathered} \text { Multi- } \\ \text { ancestry } \\ (23,948 \mathrm{ca}) \\ \mathrm{P}_{\text {random }} \end{gathered}$ | $\begin{gathered} \text { European } \\ \text { ancestry } \\ (19,954 \text { ca) } \\ \mathrm{P}_{\text {random }} \end{gathered}$ | African ancestry (2,149 ca) $\mathrm{P}_{\text {fixed }}$ | $\begin{gathered} \text { Japanese } \\ (1,239 \text { ca }) \\ \mathrm{P}_{\text {fixed }} \end{gathered}$ | Latino ( 606 ca ) P | Pediatric (8,976 ca) <br> $\mathrm{P}_{\text {random }}$ | $r^{2}$ <br> (TAGC <br> SNP, <br> catalog <br> SNP) | Catalog SNP | P-value | Pubmed.id | Mapped Genes | $\left.\begin{gathered} \text { Ancestry of } \\ \text { published } \\ \text { studies } \end{gathered} \right\rvert\,$ | Total \# cases in published study | \#cases <br> in study discovery samples | \#cases in study replication samples | \# cases from published study in TAGC | \% of study <br> cases in <br> TAGC <br> Multi- <br> ancestry | \% of <br> study cases in TAGC EU | $\begin{gathered} \text { study } \\ \text { cases } \\ \text { in } \\ \text { TAGC } \\ \text { AF } \end{gathered}$ | $\begin{gathered} \text { study } \\ \text { cases } \\ \text { in } \\ \text { TAGC } \\ \text { JAP } \end{gathered}$ | $\left.\begin{array}{\|c\|} \hline \% \text { of } \\ \text { study } \\ \text { cases in } \\ \text { TAGC } \\ \text { LAT } \end{array} \right\rvert\,$ |
| 7q22.3 | 105,658,450 | rs6967330 | $9.1 \times 10^{-4}$ | $3.2 \times 10^{-3}$ | 0.37 | 0.21 | NA | $1.7 \times 10^{-3}$ | 1 | rs6967330 | $\begin{aligned} & \mathrm{P}_{\mathrm{fix}}=3 \times 10^{-14^{* * *}} \\ & \mathrm{P}_{\mathrm{ran}}=3 \times 10^{-7} \end{aligned}$ | 24241537 | CDHR3 | $\begin{array}{\|l\|} \mathrm{EU}(\mathrm{D}) ; \\ \mathrm{EU}(\mathrm{R}) \end{array}$ | 8,414 | 1,173 | 7,241 | 6,876 | 0.29 | 0.34 | NA | NA | NA |
| 10q24.2 | 100,739,769 | rs17111026* |  | 0.02 | 0.47 | 0.87 | 0.91 | 0.72 | 0.76 | rs12570188 | $5.0 \times 10^{-8}$ | 23829686 | HPSE2 | EU, AF, LAT | 527 | one ethnic <br> group | other ethnic groups | 0 | 0 | 0 | 0 | NA | 0 |
| 4 q 12 | 59,213,845 | rs17218161* |  | NA | NA | NA | NA | NA | NA | rs17218161 | $2.0 \times 10^{-8}$ | 23829686 | SRIP1 - <br> MIR548AG1 | EU, AF, LAT | 527 | one ethnic group | other ethnic groups | 0 | 0 | 0 | 0 | NA | 0 |
| 9 p 23 | 12,521,825 | rs16929097* | * 0.81 | 0.62 | 0.31 | NA | 0.85 | 0.13 | 1 | rs16929097 | $8.0 \times 10^{-9}$ | 23829686 | JKAMPP1- <br> TYRP1 | EU, AF, LAT | 527 | one ethnic group | other ethnic groups | 0 | 0 | 0 | 0 | NA | 0 |
| 1 q 21.3 | 154,426,263 | rs4129267 | $8.5 \times 10^{-7}$ | $4.5 \times 10^{-6}$ | 0.92 | $1.5 \times 10^{-3}$ | 0.68 | 0.08 | 1 | rs4129267 | $2.0 \times 10^{-8}$ | 21907864 | IL6R | EU | 15,797 | 12,475 | 3,322 | 12,687 | 0.53 | 0.64 | NA | NA | NA |
| 5 q 12.1 | 59,369,793 | rs1588265 | 0.55 | 0.65 | 0.79 | 0.95 | 0.42 | 0.59 | 1 | rs1588265 | $3.0 \times 10^{-8}$ | 19426955 | PDE4D | EU (D); EU, AF, LAT (R) | 7,346 | 359 | 6,987 | 3,011 | 0.13 | 0.15 | 0.21 | NA | 0.58 |
| 12q13.2 | 56,412,486 | rs1701704 | $1.5 \times 10^{-3}$ | $5.2 \times 10^{-3}$ | 0.67 | $4.5 \times 10^{-4}$ | 0.22 | 0.21 | 1 | rs1701704 | $2.0 \times 10^{-13}$ | 21804548 | IKZF4 | Jap | 7,171 | 1,532 | 5,639 | 301 | 0.01 | NA | NA | 0.24 | NA |
| 4q31.21 | 144,003,158 | rs7686660 | 0.88 | 0.99 | 0.86 | 0.69 | 0.53 | 0.19 | 1 | rs7686660 | $2.0 \times 10^{-12}$ | 21804548 | LOC729675 | JAP | 7,171 | 1,532 | 5,639 | 301 | 0.01 | NA | NA | 0.24 | NA |
| 22 q 12.3 | 37,534,033 | rs2284033 | $7.9 \times 10^{-5}$ | $5.3 \times 10^{-5}$ | 0.58 | 0.49 | 0.52 | $1.6 \times 10^{-3}$ | 1 | rs2284033 | $1.0 \times 10^{-8}$ | 20860503 | IL2RB | EU | 10,365 | 10,365 | NA | 8,082 | 0.34 | 0.41 | NA | NA | NA |
| 8q24.11 | 118,025,644 | rs3019885 | 0.99 | 0.81 | 0.11 | 0.47 | 0.09 | 0.86 | 1 | rs3019885 | $5.0 \times 10^{-13}$ | 21814517 | SLC30A8 | $\left\lvert\, \begin{aligned} & \operatorname{JAP}(\mathrm{D}) ; \\ & \operatorname{JAP}, \mathrm{KOR}(\mathrm{R}) \end{aligned}\right.$ | 2,591 | 938 | 1,653 | 938 | 0.04 | NA | NA | 0.76 | NA |
| 1q31.3 | 197,325,907 | rs2786098 | 0.87 | 0.49 | 0.96 | 0.15 | 0.23 | 0.83 | 1 | rs2786098 | $2.0 \times 10^{-13}$ | 20032318 | CRB1, DENND1B | EU, AF | 3,377 | 793 | 2,584 | 1,088 | 0.05 | 0.03 | 0.21 | NA | NA |
| 1q23.1 | 158,932,554 | rs1101999** | NA | NA | $1.1 \times 10^{-4}$ | NA | NA | NA | 1 | rs1101999 | $4.0 \times 10^{-9}$ | 21804549 | PYHIN1 | $\int_{(\mathrm{R})}^{\mathrm{AF}(\mathrm{D}) ; \mathrm{AF}}$ | 3,759 | 1,612 | 2,147 | 1,257 | 0.05 | NA | 0.58 | NA | NA |

The number of cases included in each TAGC meta-analysis is shown in parentheses
EU=European-ancestry, AF=African-ancestry, JAP=Japanese, LAT= Latino, KOR=Korean; D = discovery; R =replication
*The frequency of effect allele at 10p24.2, 4q12 and 9p23 loci (Pubmed.id=23829686) is <2\% in European-ancestry populations ( $81 \%$ of the 527 study trios are from European-ancestry); no SNP found in LD with rs17218161 in TAGC
**The frequency of effect allele at 1 q 23.1 locus (Pubmed.id $=21804549$ ) is 0.30 in African ancestry populations in whom it was discovered but is very low in other populations
*** Only $\mathrm{P}_{\text {fixed }}=3 \times 10^{-14}$ is reported in the catalog while in the publication (PMID 24241537), there is significant evidence for heterogeneity ( $\mathrm{P}=0.02$ ) and $\mathrm{P}_{\text {random }}=3 \times 10^{-7}$ is not genome-wide significant

## Supplementary Table 13. Approximate conditional analysis of asthma-associated loci

Approximate conditional analysis, using the GCTA software (PMID 21167468), is based on summary meta-analysis statistics under a fixed-effects model. This analysis was only conducted in European-ancestry populations that are assumed to share a similar LD pattern Note that this analysis could not be done at the 9p24.1 locus which shows heterogeneity in SNP effect size across studies. At the 17q12-21, this analysis was restricted to the pediatric subgroup in which this locus does not show heterogeneity (see Online Methods for details).

${ }^{\text {a }}$ In the conditional SNP field, 'TOP' indicates we present the unadjusted results for the top SNP at each locus; otherwise the rsIDs of the top (and subsequent SNPs) used to adjust other SNPs are listed
${ }^{\text {b }}$ The gene where eventually the lead SNP lies is first indicated followed by the previous and next genes
${ }^{\text {'The }}$ effect allele is the allele for which the regression coefficients (betas) are computed; ${ }^{d}$ EAF= Effect allele frequency in European-ancestry populations.
${ }^{\text {e }}$ Under the heading meta-analysis results, beta, SE and P are the regression coefficient, standard error and P -values in fixed-effects meta-analysis of European ancestry populations
fUnder the heading conditional analysis results, beta, SE and P are the regression coefficient, standard error and P -values following adjustment by the SNP(s) listed in the 'Conditional SNP' field.
Note that the TOP SNP in European-ancestry meta-analysis under a fixed-effects may differ from the TOP SNP under a random-effects model (Table 1); these TOP SNPs are indicated by a star.

Supplementary Table 14. Associations between SNPs at the nine novel asthma loci and cis-gene expression in blood and lung
The eQTL databases for blood and lung were interrogated for the lead SNPs (shown in Table 1) and all SNPs having r $r^{2}$ between 0.50 and 1 with the lead SNPs.
This table shows the most significant associations between asthma SNPs and transcripts among all associations having Ptranscript $<10^{-5}$

${ }^{\text {a }} \mathrm{R}=$ reference allele $/ \mathrm{E}=$ effect allele
${ }^{\text {b }}$ eQTL databases are described in Online Methods
${ }^{\text {cBlood }}$ is whole blood; LCL = lymphoblastoid cell lines; Lung is whole lung tissue
${ }^{d} P_{\text {transcript }}$ is the $P$-value for test of association of SNP with gene transcript

Supplementary Table 15. Overlap between TAGC asthma association signals ( $P_{\text {random }}<10^{-4}$ ) and GWAS signals with diseases/traits in the GWAS catalog.
This Table has three parts: (i) autoimmune diseases are in the first part of the table; (ii) three allergic phenotypes in the second part; and (iii) the rest of the top 10 traits (in the number of hits in the catalog) are last. Asthma association signals are those having $P_{\text {random }}<10^{-4}$ in the multi-ancestry meta-analysis. For each disease and trait, we selected one GWAS SNP per chromosomal band as reported in the variable "Region" in the GWAS catalog; SNPs associated with multiple diseases in the GWAS catalog could appear in counts in multiple rows.

| Rank ${ }^{\text {a }}$ | Disease/Trait | Number of GWAS catalog entries | Number of SNPs associated with asthma at $P_{\text {random }}<10^{-4}$ in TAGC multi-ancestry meta-analysis | Enrichment $P$-value ${ }^{\text {b }}$ |
| :---: | :---: | :---: | :---: | :---: |
| Auto-immune diseases |  |  |  |  |
| 2 | Inflammatory bowel disease | 97 | 6 | $9.8 \times 10^{-16}$ |
| 4 | Crohn's disease | 89 | 9 | $6.3 \times 10^{-25}$ |
| 8 | Rheumatoid arthritis | 59 | 4 | $4.5 \times 10^{-11}$ |
| 11 | Ulcerative colitis | 54 | 4 | $3.1 \times 10^{-11}$ |
| 14 | Multiple sclerosis | 49 | 3 | $1.8 \times 10^{-8}$ |
| 23 | Systemic lupus erythematosus | 35 | 2 | $5.9 \times 10^{-6}$ |
| 25 | Type 1 diabetes | 34 | 5 | $2.7 \times 10^{-15}$ |
| 30 | Celiac disease | 29 | 3 | $3.6 \times 10^{-9}$ |
| 38 | Vitiligo | 23 | 4 | $8.8 \times 10^{-18}$ |
| Allergy-related phenotypes |  |  |  |  |
| 65 | Self-reported allergy | 16 | 12 | Highly significant |
| 67 | Atopic dermatitis | 14 | 5 | Highly significant |
| 99 | Allergic sensitization | 9 | 7 | Highly significant |
| Top 10 traits in the number of hits in the GWAS catalog |  |  |  |  |
| 1 | Height | 162 | 3 | $6.8 \times 10^{-7}$ |
| 3 | Blood metabolite levels | 96 | 2 | $4.5 \times 10^{-5}$ |
| 5 | Cholesterol, total | 66 | 2 | $2.1 \times 10^{-5}$ |
| 6 | HDL cholesterol | 64 | 2 | $2.0 \times 10^{-5}$ |
| 7 | Type 2 diabetes | 61 | 0 | 1 |
| 9 | Breast cancer | 56 | 0 | 1 |
| 10 | Prostate cancer | 56 | 0 | 1 |

${ }^{\text {a }}$ Rank of the disease or trait in the number of unique signals in the GWAS catalog (PMID: 24316577)
${ }^{\text {b }}$ The enrichment $P$-value is the binomial tail probability for observing the shown number of TAGC SNPs with $P_{\text {random }}<10^{-4}$ among the SNPS reported in the GWAS catalog for a given disease or trait.
We investigated overlap with nine auto-immune diseases, three allergic phenotypes and seven other traits with large number of associations, and a conservative Bonferroni correction implies an adjusted significance threshold for enrichment of 0.05/19 =0.003.

Supplementary Table 16. Overlap between TAGC asthma signals ( $\mathrm{P}_{\text {random }}<10^{-3}$ in multi-ancestry meta-analysis) and GWAS signals with diseases/traits in the GWAS catalog

| Information from the NHGRI GWAS catalog ${ }^{\text {a }}$ |  |  |  |  |  |  | Information from TAGC asthma meta-analysis |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Disease.Trait | Region | Chr_position (build 38) | Reported Genes | Mapped Gene | Strongest SNP -Risk Allele | P_Value | $P_{\text {fixed }}{ }^{\text {b }}$ | Prandom ${ }^{\text {b }}$ | $P_{\text {het }}{ }^{\text {c }}$ |
| Common traits (Other) | 1921.3 | 152,520,678 | LCE3E | CRCT1-LCE3E | rs499697-G | $1 \times 10^{-10}$ | $4.3 \times 10^{-4}$ | $9.6 \times 10^{-4}$ | 0.13 |
| Fibrinogen | 1921.3 | 154,453,788 | IL6R | IL6R | rs4129267-T | $6 \times 10^{-27}$ | $8.1 \times 10^{-8}$ | $8.5 \times 10^{-7}$ | 0.19 |
| C-reactive protein | 1 q 21.3 | 154,453,788 | IL6R | IL6R | rs 4129267-C | $2 \times 10^{-48}$ | $8.1 \times 10^{-8}$ | $8.5 \times 10^{-7}$ | 0.19 |
| Protein quantitative trait loci | 1 q 21.3 | 154,453,788 | IL6R | IL6R | rs4129267-? | $2 \times 10^{-57}$ | $8.1 \times 10^{-8}$ | $8.5 \times 10^{-7}$ | 0.19 |
| Celiac disease and Rheumatoid arthritis | 1 q 24.2 | 167,442,147 | CD247 | CD247 | rs864537-? | $2 \times 10^{-11}$ | $1.4 \times 10^{-4}$ | $4.6 \times 10^{-4}$ | 0.30 |
| Ulcerative colitis | 2 q 11.2 | 102,047,167 | IL1R2 | IL1R2-IL1R1 | rs2310173-T | $3 \times 10^{-12}$ | $2.5 \times 10^{-5}$ | $2.5 \times 10^{-5}$ | 1.00 |
| Self-reported allergy | 2 q 12.1 | 102,263,004 | IL1RL2, IL1RL1 | IL1RL2 - ILIRL1 | rs10189629-A | $2 \times 10^{-16}$ | $3.2 \times 10^{-17}$ | $7.1 \times 10^{-13}$ | 0.06 |
| Serum protein levels (sST2) | 2q12.1 | 102,316,102 | IL1RL1 | IL1RL1 | rs950880-A | $7 \times 10^{-94}$ | $8.3 \times 10^{-19}$ | $2.6 \times 10^{-17}$ | 0.41 |
| Eosinophil counts | 2 q 12.1 | 102,341,256 | IL1RL1 | IL1RL1 | rs1420101-A | $5 \times 10^{-14}$ | $3.9 \times 10^{-21}$ | $3.9 \times 10^{-21}$ | 0.61 |
| Crohn's disease | 2 q 33.1 | 198,032,171 | PLCL1 | PLCL1 | rs6738825-A | $4 \times 10^{-9}$ | $5.0 \times 10^{-4}$ | $5.0 \times 10^{-4}$ | 0.53 |
| Self-reported allergy | 2 q 33.1 | 198,049,348 | PLCL1 | PLCL1 | rs10497813-G | $6 \times 10^{-10}$ | $3.7 \times 10^{-4}$ | $3.7 \times 10^{-4}$ | 0.65 |
| Menarche (age at onset) | 3q22.1 | 132,891,908 | TMEM108, NPHP3 | NPHP3-AS1-TMEM108 | rs6439371-G | $1 \times 10^{-8}$ | $7.5 \times 10^{-4}$ | $7.5 \times 10^{-4}$ | 0.65 |
| Allergic sensitization | 3 q 28 | 188,354,725 | LPP, BCL6 | LPP | rs9865818-G | $3 \times 10^{-10}$ | $5.2 \times 10^{-5}$ | $1.6 \times 10^{-4}$ | 0.01 |
| Vitiligo | 3 q 28 | 188,394,766 | LPP | LPP | rs1464510-T | $1 \times 10^{-11}$ | $1.0 \times 10^{-6}$ | $2.2 \times 10^{-5}$ | $4.4 \times 10^{-3}$ |
| Celiac disease | 3 q 28 | 188,394,766 | LPP | LPP | rs1464510-A | $3 \times 10^{-40}$ | $1.0 \times 10^{-6}$ | $2.2 \times 10^{-5}$ | $4.4 \times 10^{-3}$ |
| Self-reported allergy | 3 q 28 | 188,411,191 | LPP, BCL6 | LPP | rs9860547-A | $1 \times 10^{-9}$ | $7.4 \times 10^{-5}$ | $3.2 \times 10^{-4}$ | $4.4 \times 10^{-3}$ |
| Self-reported allergy | 4 p 14 | 38,809,930 | TLR1,TLR6,TLR10 | TLR1-TLR6 | rs2101521-A | $5 \times 10^{-21}$ | $8.1 \times 10^{-4}$ | $8.1 \times 10^{-4}$ | 0.74 |
| Allergic sensitization | 4 p 14 | 38,811,255 | TLR1,TLR6, TLR10, MIR574, FAM114A1 | TLR1- TLR6 | rs17616434-T | $5 \times 10^{-11}$ | $1.0 \times 10^{-5}$ | $5.2 \times 10^{-5}$ | 0.22 |
| Type 1 diabetes | 4 q 27 | 122,211,337 | IL2 | KIAA1109 | rs4505848-? | $5 \times 10^{-13}$ | $2.7 \times 10^{-6}$ | $5.4 \times 10^{-5}$ | 0.21 |
| Self-reported allergy | 4 q 27 | 122,408,207 | ADAD1 | ADAD1 | rs17388568-A | $4 \times 10^{-8}$ | $4.1 \times 10^{-5}$ | $6.2 \times 10^{-4}$ | 0.11 |
| Allergic sensitization | 5 q 22.1 | 110,854,353 | SLC25A46, TSLP, WDR36, CAMK4 | SLC25A46-TSLP | rs10056340-G | $5 \times 10^{-14}$ | $3.2 \times 10^{-8}$ | $3.2 \times 10^{-8}$ | 0.67 |
| Eosinophilic esophagitis (pediatric) | 5q22.1 | 111,069,977 | WDR36 | TSLP | rs3806932-? | $3 \times 10^{-9}$ | $3.6 \times 10^{-9}$ | $4.5 \times 10^{-8}$ | 0.36 |
| Self-reported allergy | 5 q 22.1 | 111,131,801 | WDR36, CAMK4, TSLP | WDR36-RPS3AP21 | rs1438673-C | $2 \times 10^{-20}$ | $6.1 \times 10^{-19}$ | $4.2 \times 10^{-13}$ | 0.10 |
| Inflammatory bowel disease | 5 q 23.3 | 130,681,594 | Intergenic | ARL2BPP4-RPL11P2 | rs4836519-T | $4 \times 10^{-10}$ | $7.6 \times 10^{-4}$ | $7.6 \times 10^{-4}$ | 0.73 |
| Crohn's disease | 5 q 31.1 | 132,067,045 | IL3, ACSL6, P4HA2, PDLIM4, SLC22A4 | IL3-CSF2 | rs3091338-T | $4 \times 10^{-8}$ | $1.8 \times 10^{-4}$ | $3.0 \times 10^{-4}$ | 0.39 |
| Blood metabolite levels | 5 q 31.1 | 132,316,836 | SLC22A4 | SLC22A4;LOC553103 | rs11950562-A | $2 \times 10^{-41}$ | $9.0 \times 10^{-11}$ | $9.0 \times 10^{-11}$ | 0.70 |
| Blood metabolite ratios | 5 q 31.1 | 132,329,685 | SLC22A4 | SLC22A4;LOC553103 | rs272889-A | $3 \times 10^{-51}$ | $1.9 \times 10^{-5}$ | $1.9 \times 10^{-5}$ | 0.55 |
| Metabolic traits | 5 q 31.1 | 132,329,685 | SLC22A4 | SLC22A4;LOC553103 | rs272889-A | $7 \times 10^{-16}$ | $1.9 \times 10^{-5}$ | $1.9 \times 10^{-5}$ | 0.55 |
| Height | 5 q 31.1 | 132,350,453 | FU44796 | LOC553103 | rs10058074-A | $4 \times 10^{-12}$ | $2.2 \times 10^{-9}$ | $2.2 \times 10^{-9}$ | 0.69 |
| Inflammatory bowel disease | $5 q 31.1$ | 132,435,113 | IRF1, IL13, CSF2, SLC22A4, IL4, IL3, IL5, PDLIM4, SLC22A5, ACSL6 | C5orf56 | rs2188962-T | $1 \times 10^{-52}$ | $7.3 \times 10^{-10}$ | $7.1 \times 10^{-8}$ | 0.15 |
| Platelet counts | 5 q 31.1 | 132,484,229 | IRF1 | IRF1 | rs2070729-A | $1 \times 10^{-10}$ | $2.3 \times 10^{-9}$ | $4.5 \times 10^{-8}$ | 0.30 |
| Atopic dermatitis | 5 q 31.1 | 132,660,151 | IL13, RAD50 | IL13 | rs1295686-A | $2 \times 10^{-17}$ | $2.8 \times 10^{-15}$ | $2.8 \times 10^{-15}$ | 0.67 |
| lgE levels | 5 q 31.1 | 132,660,272 | IL13 | IL13 | rs20541-A | $3 \times 10^{-18}$ | $5.0 \times 10^{-16}$ | $5.0 \times 10^{-16}$ | 0.77 |
| Psoriasis | 5 q 31.1 | 132,660,272 | 1 L 13 | 1 L 13 | rs20541-G | $5 \times 10^{-15}$ | $5.0 \times 10^{-16}$ | $5.0 \times 10^{-16}$ | 0.77 |
| Hodgkin's lymphoma | 5 q 31.1 | 132,662,721 | IL13 | IL13-1L4 | rs2069757-A | $2 \times 10^{-11}$ | $3.3 \times 10^{-7}$ | $3.3 \times 10^{-7}$ | 0.82 |
| Crohn's disease | 5 q 31.3 | 142,099,500 | NDFIP1 | MRPL11P2 - NDFIP1 | rs11167764-C | $2 \times 10^{-9}$ | $1.5 \times 10^{-8}$ | $1.5 \times 10^{-6}$ | 0.12 |
| Schizophrenia or bipolar disorder | 6 p 22.1 | 27,742,386 | MHC | TRNAI25 | rs17693963-? | $3 \times 10^{-11}$ | $6.0 \times 10^{-7}$ | $6.0 \times 10^{-7}$ | 0.74 |
| Schizophrenia, schizoaffective disorder or bipolar disorder | 6 p 22.1 | 27,742,386 | MHC | TRNAI25 | rs17693963-? | $2 \times 10^{-9}$ | $6.0 \times 10^{-7}$ | $6.0 \times 10^{-7}$ | 0.74 |
| Pulmonary function | 6 p 22.1 | 28,354,519 | ZKSCAN3, ZNF323 | ZKSCAN3 | rs6903823-G | $2 \times 10^{-10}$ | $8.6 \times 10^{-8}$ | $8.6 \times 10^{-8}$ | 0.62 |
| Barrett's esophagus | 6 p 22.1 | 29,388,554 | MHC, OR2D12, OR2D13 | TRNA125 | rs9257809-A | $4 \times 10^{-9}$ | $3.8 \times 10^{-6}$ | $3.8 \times 10^{-6}$ | 0.77 |
| Autism spectrum disorder, attention deficithyperactivity disorder, bipolar disorder, major depressive disorder, and schizophrenia (combined) | 6 p 22.1 | 30,064,745 | MHC region | PPP1R11;ZNRD1 | rs8321-? | $8 \times 10^{-9}$ | $4.6 \times 10^{-7}$ | $4.6 \times 10^{-7}$ | 0.93 |
| Schizophrenia | 6 p 22.1 | 30,197,496 | MHC, TRIM26 | TRIM26 | rs2523722-G | $1 \times 10^{-16}$ | $5.7 \times 10^{-8}$ | $5.7 \times 10^{-8}$ | 0.79 |
| Myasthenia gravis | 6 p 21.33 | 31,090,563 | NR | TRNA125 | rs3130544-A | $2 \times 10^{-90}$ | $1.8 \times 10^{-8}$ | $1.8 \times 10^{-8}$ | 0.48 |
| Chronic hepatitis B infection | 6 p 21.33 | 31,162,816 | TCF19 | TCF19 | rs1419881-? | $1 \times 10^{-18}$ | $6.9 \times 10^{-5}$ | $1.2 \times 10^{-4}$ | 0.30 |
| HIV-1 control | 6 p 21.33 | 31,175,805 | PSORS1C3 | PSORS1C3 | rs3131018-C | $4 \times 10^{-16}$ | $1.4 \times 10^{-4}$ | $2.3 \times 10^{-4}$ | 0.35 |
| Psoriasis | 6p21.33 | 31,344,549 | HLA-C | TRNAI25 | rs3134792-? | $1 \times 10^{-9}$ | $3.1 \times 10^{-6}$ | $5.0 \times 10^{-6}$ | 0.44 |


| Information from the NHGRI GWAS catalog ${ }^{\text {a }}$ |  |  |  |  |  |  | Information from TAGC asthma meta-analysis |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Disease.Trait | Region | Chr_position (build 38) | Reported Genes | Mapped Gene | Strongest SNP -Risk Allele | P_Value | $P_{\text {fixed }}{ }^{\text {b }}$ | $P_{\text {random }}{ }^{\text {b }}$ | $P_{\text {het }}{ }^{\text {c }}$ |
| Self-reported allergy | 6p21.33 | 31,384,336 | HLA-C, MICA | TRNA125 | rs9266772-C | $3 \times 10^{-12}$ | $8.5 \times 10^{-6}$ | $2.3 \times 10^{-5}$ | 0.36 |
| Rheumatoid arthritis (ACPA-negative) | 6p21.33 | 31,385,552 | HLA-B,MICA | TRNAI25 | rs2596565-? | $9 \times 10^{-9}$ | $1.0 \times 10^{-7}$ | $1.0 \times 10^{-6}$ | 0.27 |
| Allergic sensitization | 6p21.33 | 31,386,405 | HLA-B, MICA, HLA-C, MICB | TRNAI25 | rs6932730-T | $4 \times 10^{-8}$ | $5.2 \times 10^{-5}$ | $2.0 \times 10^{-4}$ | 0.31 |
| Height | 6p21.33 | 31,386,783 | HLA-B | TRNAI25 | rs13437082-? | $5 \times 10^{-8}$ | $5.8 \times 10^{-5}$ | $2.9 \times 10^{-4}$ | 0.11 |
| Beta-2 microglubulin plasma levels | 6 p 21.33 | 31,394,533 | HLA-B | LOC101929072 | rs16899524-C | $1 \times 10^{-8}$ | $7.5 \times 10^{-6}$ | $1.0 \times 10^{-5}$ | 0.44 |
| Metabolic syndrome | 6 p 21.33 | 31,481,199 | HCG26, MICB | TRNAI25 | rs3099844-A | $2 \times 10^{-8}$ | $1.4 \times 10^{-9}$ | $1.4 \times 10^{-9}$ | 0.56 |
| Neonatal lupus | 6p21.33 | 31,481,199 | TNF,NFKBIL1,LTA,LTB,AIF1 | TRNA125 | rs3099844-? | $5 \times 10^{-10}$ | $1.4 \times 10^{-9}$ | $1.4 \times 10^{-9}$ | 0.56 |
| Dengue shock syndrome | 6p21.33 | 31,507,709 | MICB | MICB | rs3132468-? | $4 \times 10^{-11}$ | $1.4 \times 10^{-7}$ | $1.2 \times 10^{-5}$ | 0.16 |
| Stevens-Johnson syndrome and toxic epidermal necrolysis (SJS-TEN) | 6p21.33 | 31,537,703 | BAT1, HCP5, MICC, PPIAP9, PSORS1C1, POLR2LP, CCHCR1, TCF19, POU5F1, HLA-C, HCP5, PSORS1C3, HLA-B*5801 | DDX39B;SNORD117;ATP6V1G2- DDX39B | rs2734583-? | $2 \times 10^{-8}$ | $3.8 \times 10^{-8}$ | $3.8 \times 10^{-8}$ | 0.65 |
| Lung adenocarcinoma | 6 p 21.33 | 31,652,743 | BAT3, APOM | BAG6;APOM | rs3117582-C | $5 \times 10^{-12}$ | $7.1 \times 10^{-7}$ | $7.1 \times 10^{-7}$ | 0.51 |
| Lung cancer | 6p21.33 | 31,652,743 | BAT3 | BAG6;APOM | rs3117582-? | $4 \times 10^{-10}$ | $7.1 \times 10^{-7}$ | $7.1 \times 10^{-7}$ | 0.51 |
| Systemic lupus erythematosus | 6 p 21.33 | 31,902,549 | C2 | C2;ZBTB12 | rs558702-A | $8 \times 10^{-21}$ | $1.8 \times 10^{-6}$ | $1.8 \times 10^{-6}$ | 0.51 |
| Hematology traits | 6p21.33 | 31,973,120 | ZBTB12, CFB, STK19 | DXO;STK19 | rs389884-C | $2 \times 10^{-8}$ | $1.4 \times 10^{-6}$ | $1.4 \times 10^{-6}$ | 0.56 |
| Serum total protein level | 6 p 21.32 | 32,142,202 | Intergenic | TRNAI25 | rs204999-A | $3 \times 10^{-9}$ | $2.3 \times 10^{-4}$ | $5.6 \times 10^{-4}$ | 0.01 |
| Phospholipid levels (plasma) | 6 p 21.32 | 32,168,770 | AGPAT1 | AGPAT1;EGFL8;PPT2-EGFL8 | rs1061808-? | $8 \times 10^{-10}$ | $4.9 \times 10^{-9}$ | $4.9 \times 10^{-9}$ | 0.56 |
| Atopic dermatitis | 6 p 21.32 | 32,190,542 | GPSM3 | PBX2;GPSM3 | rs176095-T | $8 \times 10^{-20}$ | $4.8 \times 10^{-5}$ | $2.2 \times 10^{-4}$ | $9.0 \times 10^{-3}$ |
| Complement C3 and C4 levels | 6p21.32 | 32,197,667 | HLA-DRA | NOTCH4 | rs2071278-G | $4 \times 10^{-72}$ | $4.6 \times 10^{-5}$ | $3.5 \times 10^{-4}$ | 0.10 |
| Systemic sclerosis | 6 p 21.32 | 32,222,629 | NOTCH4 | NOTCH4 | rs 443198-? | $9 \times 10^{-21}$ | $3.2 \times 10^{-6}$ | $1.7 \times 10^{-5}$ | 0.26 |
| Nephropathy | 6 p 21.32 | 32,251,066 | Intergenic | C6orf10 | rs3115573-? | $1 \times 10^{-9}$ | $5.8 \times 10^{-5}$ | $5.8 \times 10^{-5}$ | 0.55 |
| Chronic lymphocytic leukemia | 6 p 21.32 | 32,289,789 | HLA | C6orf10 | rs926070-A | $4 \times 10^{-8}$ | $1.2 \times 10^{-6}$ | $1.2 \times 10^{-6}$ | 0.51 |
| Lumiracoxib-related liver injury | 6 p 21.32 | 32,338,202 | HLA-DRB1 | C6orf10 | rs3129900-? | $7 \times 10^{-25}$ | $1.4 \times 10^{-5}$ | $5.6 \times 10^{-5}$ | 0.32 |
| Multiple sclerosis | 6p21.32 | 32,368,410 | C6orf10 | C6orf10 | rs3129934-T | $7 \times 10^{-16}$ | $1.2 \times 10^{-5}$ | $5.9 \times 10^{-5}$ | 0.30 |
| Graves' disease | 6 p 21.32 | 32,369,853 | MHC | C6orf10 | rs2273017-A | $2 \times 10^{-22}$ | $1.0 \times 10^{-5}$ | $1.0 \times 10^{-5}$ | 0.68 |
| Hepatitis C induced liver cirrhosis | 6 p 21.32 | 32,400,310 | BTNL2 | BTNL2 | rs3817963-A | $1 \times 10^{-8}$ | $2.7 \times 10^{-19}$ | $6.4 \times 10^{-13}$ | 0.06 |
| Lung adenocarcinoma | 6 p 21.32 | 32,400,310 | BTNL2 | BTNL2 | rs3817963-G | $3 \times 10^{-10}$ | $2.7 \times 10^{-19}$ | $6.4 \times 10^{-13}$ | 0.06 |
| Multiple sclerosis (OCB status) | 6 p 21.32 | 32,400,310 | BTNL2 | BTNL2 | rs3817963-? | $6 \times 10^{-10}$ | $2.7 \times 10^{-19}$ | $6.4 \times 10^{-13}$ | 0.06 |
| Vitiligo | 6 p 21.32 | 32,405,921 | BTNL2, HLA-DRA, HLA-DQA1 | BTNL2;LOC101929163 | rs3806156-T | $7 \times 10^{-19}$ | $6.3 \times 10^{-11}$ | $1.1 \times 10^{-6}$ | $1.5 \times 10^{-3}$ |
| Parkinson's disease | 6 p 21.32 | 32,420,032 | LOC642072 | TRNAI25 | rs2395163-? | $3 \times 10^{-11}$ | $5.8 \times 10^{-17}$ | $1.5 \times 10^{-10}$ | 0.09 |
| Cholesterol, total | 6 p 21.32 | 32,444,658 | HLA | HLA-DRA | rs3177928-A | $1 \times 10^{-21}$ | $1.3 \times 10^{-6}$ | $2.0 \times 10^{-5}$ | 0.16 |
| LDL cholesterol | 6p21.32 | 32,444,658 | HLA | HLA-DRA | rs3177928-A | $3 \times 10^{-17}$ | $1.3 \times 10^{-6}$ | $2.0 \times 10^{-5}$ | 0.16 |
| Hodgkin's lymphoma | 6 p 21.32 | 32,460,508 | HLA class II, HLA class I | TRNAI25 | rs6903608-C | $7 \times 10^{-31}$ | $5.4 \times 10^{-5}$ | $3.8 \times 10^{-4}$ | 0.20 |
| Rheumatoid arthritis | 6 p 21.32 | 32,461,866 | HLA-DRA | TRNAI25 | rs9268853-C | $5 \times 10^{-10} 9$ | $1.8 \times 10^{-18}$ | $6.1 \times 10^{-17}$ | 0.39 |
| Ulcerative colitis | 6p21.32 | 32,461,866 | HLA-DRB5, HLA-DQA1, HLA-DRB1, HLA-DRA, BTNL2 | TRNAI25 | rs9268853-T | $1 \times 10^{-55}$ | $1.8 \times 10^{-18}$ | $6.1 \times 10^{-17}$ | 0.39 |
| $\lg A$ nephropathy | 6p21.32 | 32,609,603 | HLA-DRB1 | TRNAI25 | rs660895-G | $4 \times 10^{-20}$ | $3.2 \times 10^{-16}$ | $1.1 \times 10^{-6}$ | $8.0 \times 10^{-4}$ |
| Lymphoma | 6 p 21.32 | 32,614,112 | HLA-DRB5, HLA-DQA1 | TRNAI25 | rs4530903-T | $2 \times 10^{-8}$ | $1.1 \times 10^{-8}$ | $2.0 \times 10^{-7}$ | 0.13 |
| Schizophrenia | 6 p 21.32 | 32,634,492 | HLA-DQA1 | TRNAI25 | rs9272219-G | $7 \times 10^{-8}$ | $2.0 \times 10^{-5}$ | $2.9 \times 10^{-5}$ | 0.42 |
| Type 1 diabetes | 6p21.32 | 32,636,595 | HLA | HLA-DQA1 | rs9272346-G | $6 \times 10^{-12} 9$ | $8.2 \times 10^{-32}$ | $5.7 \times 10^{-24}$ | 0.14 |
| Immunoglobulin A | 6 p 21.32 | 32,638,107 | HLA-DRB1 | HLA-DQA1 | rs2187668-A | $2 \times 10^{-33}$ | $1.0 \times 10^{-5}$ | $5.3 \times 10^{-5}$ | 0.31 |
| Celiac disease | 6 p 21.32 | 32,638,107 | HLA-DQA1, HLA-DQB1 | HLA-DQA1 | rs2187668-A | $1 \times 10^{-50}$ | $1.0 \times 10^{-5}$ | $5.3 \times 10^{-5}$ | 0.31 |
| Autoimmune hepatitis type-1 | 6 p 21.32 | 32,638,107 | HLA-DQA1 | HLA-DQA1 | rs2187668-? | $2 \times 10^{-78}$ | $1.0 \times 10^{-5}$ | $5.3 \times 10^{-5}$ | 0.31 |
| Nephropathy (idiopathic membranous) | 6 p 21.32 | 32,638,107 | HLA-DQA1 | HLA-DQA1 | rs2187668-? | $8 \times 10^{-93}$ | $1.0 \times 10^{-5}$ | $5.3 \times 10^{-5}$ | 0.31 |
| Allergic sensitization | 6p21.32 | 32,658,534 | HLA-DQB1, HLA-DQA1, | TRNAI25 | rs6906021-C | $2 \times 10^{-12}$ | $3.9 \times 10^{-14}$ | $8.4 \times 10^{-10}$ | 0.05 |
| Self-reported allergy | 6 p 21.32 | 32,658,534 | HLA-DQA1, HLA-DQB1 | TRNAI25 | rs6906021-C | $7 \times 10^{-15}$ | $3.9 \times 10^{-14}$ | $8.4 \times 10^{-10}$ | 0.05 |
| Follicular lymphoma | 6 p 21.32 | 32,696,681 | HLA-DQB1 | TRNAI25 | rs2647012-G | $2 \times 10^{-21}$ | $1.0 \times 10^{-6}$ | $6.7 \times 10^{-4}$ | 0.02 |
| HPV seropositivity | 6 p 21.32 | 32,697,183 | HLA-DQB1 | TRNAI25 | rs9357152-G | $1 \times 10^{-14}$ | $1.6 \times 10^{-7}$ | $4.5 \times 10^{-7}$ | 0.41 |
| Hepatocellular carcinoma (hepatitis $B$ virus related) | 6p21.32 | 32,698,518 | HLA-DQ | TRNAI25 | rs9275319-A | $3 \times 10^{-17}$ | $7.6 \times 10^{-5}$ | $6.0 \times 10^{-4}$ | 0.31 |
| Hepatitis B | 6 p 21.32 | 32,702,478 | HLA-DQB1 | TRNAI25 | rs2856718-A | $4 \times 10^{-37}$ | $2.0 \times 10^{-8}$ | $1.9 \times 10^{-7}$ | 0.27 |
| Chronic hepatitis B infection | 6 p 21.32 | 32,702,478 | HLA-DQ | TRNAI25 | rs2856718-? | $2 \times 10^{-24}$ | $2.0 \times 10^{-8}$ | $1.9 \times 10^{-7}$ | 0.27 |
| Systemic lupus erythematosus | 6 p 21.32 | 32,711,222 | HLA-DQA2 | TRNA125 | rs9275572-A | $5 \times 10^{-16}$ | $4.8 \times 10^{-7}$ | $4.1 \times 10^{-4}$ | 0.04 |
| Chronic hepatitis C infection | 6 p 21.32 | 32,711,222 | HLA-DQB1, HLA-DQA1 | TRNAI25 | rs9275572-T | $4 \times 10^{-16}$ | $4.8 \times 10^{-7}$ | $4.1 \times 10^{-4}$ | 0.04 |
| Hepatocellular carcinoma | 6 p 21.32 | 32,711,222 | HLA-DQ, HLA-DR | TRNAI25 | rs9275572-A | $6 \times 10^{-9}$ | $4.8 \times 10^{-7}$ | $4.1 \times 10^{-4}$ | 0.04 |
| Alopecia areata | 6 p 21.32 | 32,711,222 | HLA-DQA2 | TRNAI25 | rs9275572-G | $1 \times 10^{-35}$ | $4.8 \times 10^{-7}$ | $4.1 \times 10^{-4}$ | 0.04 |


| Information from the NHGRI GWAS catalog ${ }^{\text {a }}$ |  |  |  |  |  |  | Information from TAGC asthma meta-analysis |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Disease.Trait | Region | Chr_position (build 38) | Reported Genes | Mapped Gene | Strongest SNP -Risk Allele | P_Value | $P_{\text {fixed }}{ }^{\text {b }}$ | Prandom ${ }^{\text {b }}$ | $P_{\text {het }}{ }^{\text {c }}$ |
| Pulmonary function (interaction) | 6 p 21.32 | 32,712,799 | HLA-DQB1, HLA-DQA2 | TRNA125 | rs7764819-T | $4 \times 10^{-9}$ | $1.2 \times 10^{-5}$ | $1.2 \times 10^{-5}$ | 0.50 |
| Crohn's disease | 6p21.32 | 32,713,151 | HLA-DQA2, HLA-DRB1, HLA-DQA1, HLA-DQB1, HLA-DOB, PSMB9 | TRNAI25 | rs7765379-G | $9 \times 10^{-59}$ | $8.2 \times 10^{-6}$ | $8.2 \times 10^{-6}$ | 0.55 |
| Thyroid peroxidase antibody positivity | $6 q 15$ | 90,170,674 | BACH2 | BACH2 | rs10944479-A | $4 \times 10^{-8}$ | $2.2 \times 10^{-6}$ | $2.2 \times 10^{-6}$ | 0.93 |
| Celiac disease | $6 q 15$ | 90,216,893 | BACH2, MAP3K7 | BACH2 | rs10806425-A | $4 \times 10^{-10}$ | $3.6 \times 10^{-6}$ | $3.6 \times 10^{-6}$ | 0.85 |
| Vitiligo | 6 q 15 | 90,247,744 | BACH2 | BACH2 | rs3757247-A | $3 \times 10^{-8}$ | $4.2 \times 10^{-6}$ | $4.2 \times 10^{-6}$ | 0.79 |
| Inflammatory bowel disease | 6 q 15 | 90,263,440 | Intergenic | BACH2 | rs1847472-C | $2 \times 10^{-10}$ | $1.4 \times 10^{-11}$ | $1.4 \times 10^{-11}$ | 0.74 |
| Crohn's disease | 6 q 15 | 90,263,440 | BACH2 | BACH2 | rs1847472-G | $5 \times 10^{-9}$ | $1.4 \times 10^{-11}$ | $1.4 \times 10^{-11}$ | 0.74 |
| Multiple sclerosis | 6 q 15 | 90,287,050 | BACH2 | BACH2 | rs12212193-G | $4 \times 10^{-8}$ | $6.7 \times 10^{-11}$ | $6.7 \times 10^{-11}$ | 0.58 |
| Rheumatoid arthritis | 6 q 21 | 106,219,660 | ATG5 | ATG5 | rs9372120-G | $8 \times 10^{-10}$ | $1.1 \times 10^{-5}$ | $1.1 \times 10^{-5}$ | 0.48 |
| Pulmonary function | 6 q 21 | 108,946,847 | ARMC2 | ARMC2 | rs2798641-T | $8 \times 10^{-9}$ | $9.6 \times 10^{-4}$ | $9.6 \times 10^{-4}$ | 0.74 |
| Multiple sclerosis | 6 q 23.3 | 137,646,077 | OLIG3 | BTF3L4P3-TNFAIP3 | rs13192841-A | $1 \times 10^{-8}$ | $9.5 \times 10^{-5}$ | $3.5 \times 10^{-4}$ | 0.02 |
| Mean corpuscular hemoglobin | 6 q 24.1 | 139,514,552 | CITED2 | CITED2 - ATP5F1P6 | rs668459-T | $9 \times 10^{-9}$ | $2.3 \times 10^{-5}$ | $8.5 \times 10^{-4}$ | 0.01 |
| Mean corpuscular volume | 6q24.1 | 139,514,552 | CITED2 | CITED2 - ATP5F1P6 | rs668459-T | $4 \times 10^{-8}$ | $2.3 \times 10^{-5}$ | $8.5 \times 10^{-4}$ | 0.01 |
| Atopic dermatitis | 7p22.2 | 3,089,155 | CARD11 | CARD11-SDK1 | rs $4722404-\mathrm{G}$ | $8 \times 10^{-9}$ | $1.5 \times 10^{-5}$ | $1.5 \times 10^{-5}$ | 0.48 |
| QRS duration | 7p12.3 | 46,580,547 | IGFBP3 | TTC4P1-HMGN1P19 | rs7784776-G | $1 \times 10^{-9}$ | $5.6 \times 10^{-4}$ | $5.6 \times 10^{-4}$ | 0.58 |
| Self-reported allergy | 8q21.13 | 80,355,920 | TPD52, ZBTB10 | RPS5P5-ZBTB10 | rs6473223-T | $8 \times 10^{-8}$ | $1.3 \times 10^{-10}$ | $1.3 \times 10^{-10}$ | 0.49 |
| Self-reported allergy | 9 p 24.1 | 6,172,380 | RANBP6, IL33 | RANBP6-GTF3AP1 | rs7032572-G | $2 \times 10^{-9}$ | $1.3 \times 10^{-24}$ | $1.4 \times 10^{-16}$ | 0.06 |
| Coronary heart disease | 9 p 21.3 | 22,103,814 | CDKN2A/2B | CDKN2B-AS1 | rs1333042-? | $1 \times 10^{-9}$ | $5.1 \times 10^{-4}$ | $5.1 \times 10^{-4}$ | 0.82 |
| Breast cancer | 9 q 31.2 | 108,126,198 | Intergenic | CHCHD4P2-RPL36P14 | rs865686-T | $1 \times 10^{-34}$ | $2.8 \times 10^{-4}$ | $7.1 \times 10^{-4}$ | 0.39 |
| Blood metabolite levels | 9 q 3.2 | 133,278,431 | ABO | ABO-SURF6 | rs651007-T | $6 \times 10^{-20}$ | $4.4 \times 10^{-4}$ | $4.4 \times 10^{-4}$ | 0.99 |
| Serum alkaline phosphatase levels | 9 q 3.2 | 133,278,431 | SURF6, ABO | ABO-SURF6 | rs651007-A | $1 \times 10^{-56}$ | $4.4 \times 10^{-4}$ | $4.4 \times 10^{-4}$ | 0.99 |
| End-stage coagulation | 9 q 34.2 | 133,278,431 | ABO | ABO-SURF6 | rs651007-C | $2 \times 10^{-25}$ | $4.4 \times 10^{-4}$ | $4.4 \times 10^{-4}$ | 0.99 |
| Metabolite levels | 9 q 34.2 | 133,278,431 | ABO | ABO-SURF6 | rs651007-A | $6 \times 10^{-9}$ | $4.4 \times 10^{-4}$ | $4.4 \times 10^{-4}$ | 0.99 |
| E -selectin levels | 9 q 3.2 | 133,278,431 | ABO | ABO-SURF6 | rs651007-T | $2 \times 10^{-82}$ | $4.4 \times 10^{-4}$ | $4.4 \times 10^{-4}$ | 0.99 |
| Urinary metabolites (H-NMR features) | 9 q 34.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-T | $1 \times 10^{-28}$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Red blood cell traits | 9 q 34.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-T | $9 \times 10^{-18}$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Liver enzyme levels (alkaline phosphatase) | 9 q 34.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-T | $3 \times 10^{-12} 3$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Coronary heart disease | 9 q 3.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-C | $4 \times 10^{-14}$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Soluble levels of adhesion molecules | 9 q 34.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-T | $2 \times 10^{-41}$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Soluble E-selectin levels | 9 q 3.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-C | $1 \times 10^{-29}$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Coronary artery disease or ischemic stroke | 9 q 3.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-? | $2 \times 10^{-9}$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Coronary artery disease or large artery stroke | 9 q 3.2 | 133,278,724 | ABO | ABO-SURF6 | rs579459-? | $3 \times 10^{-8}$ | $3.1 \times 10^{-4}$ | $3.1 \times 10^{-4}$ | 1.00 |
| Blood metabolite ratios | 9 q 34.2 | 133,278,860 | ABO | ABO-SURF6 | rs649129-T | $9 \times 10^{-37}$ | $2.9 \times 10^{-4}$ | $2.9 \times 10^{-4}$ | 1.00 |
| Venous thromboembolism | 9 q 3.2 | 133,279,294 | ABO | ABO-SURF6 | rs 495828-T | $3 \times 10^{-16}$ | $2.6 \times 10^{-4}$ | $2.6 \times 10^{-4}$ | 1.00 |
| Hematological and biochemical traits | 9q34.2 | 133,279,294 | ABO | ABO-SURF6 | rs 495828 -T | $4 \times 10^{-59}$ | $2.6 \times 10^{-4}$ | $2.6 \times 10^{-4}$ | 1.00 |
| Red blood cell count | 9 q 34.2 | 133,279,294 | ABO | ABO-SURF6 | rs 495828-T | $3 \times 10^{-12}$ | $2.6 \times 10^{-4}$ | $2.6 \times 10^{-4}$ | 1.00 |
| Angiotensin-converting enzyme activity | 9 q 3.2 | 133,279,294 | ABO | ABO-SURF6 | rs495828-A | $3 \times 10^{-8}$ | $2.6 \times 10^{-4}$ | $2.6 \times 10^{-4}$ | 1.00 |
| Self-reported allergy | 10p14 | 9,011,169 | GATA3 | RNA5SP299-LINC00709 | rs962993-T | $2 \times 10^{-8}$ | $6.7 \times 10^{-9}$ | $3.3 \times 10^{-6}$ | 0.13 |
| Breast cancer | 10p12.31 | 21,744,013 | MLLT10, DNAJC1 | MLLT10 | rs7072776-A | $4 \times 10^{-14}$ | $1.2 \times 10^{-4}$ | $1.2 \times 10^{-4}$ | 0.77 |
| Crohn's disease | 10q21.2 | 62,710,915 | Intergenic | ZNF365-ALDH7A1P4 | rs224136-? | $1 \times 10^{-10}$ | $6.9 \times 10^{-5}$ | $2.5 \times 10^{-4}$ | 0.28 |
| Mean platelet volume | 10q21.3 | 63,290,899 | JMJD1C, NRBF2, REEP3 | JMJD1C | rs7075195-A | $3 \times 10^{-18}$ | $2.3 \times 10^{-4}$ | $2.9 \times 10^{-4}$ | 0.45 |
| Forced vital capacity | 11p11.2 | 45,229,181 | PRDM11 | PRDM11 | rs2863171-C | $9 \times 10^{-10}$ | $3.8 \times 10^{-4}$ | $3.8 \times 10^{-4}$ | 0.51 |
| Blood metabolite levels | 11q12.2 | 61,783,884 | FADS1 | MYRF | rs174535-T | $2 \times 10^{-94}$ | $2.9 \times 10^{-5}$ | $2.9 \times 10^{-5}$ | 1.00 |
| Phospholipid levels (plasma) | 11912.2 | 61,784,455 | C11orf9 | MYRF | rs174536-A | $1 \times 10^{-63}$ | $1.8 \times 10^{-5}$ | $1.8 \times 10^{-5}$ | 1.00 |
| Oleic acid (18:1n-9) plasma levels | 11912.2 | 61,790,331 | C11orf10, C11orf9, FADS1, FADS2, FADS3, FEN1, RAB3IL1 | TMEM258 | rs102275-C | $2 \times 10^{-32}$ | $3.8 \times 10^{-5}$ | $3.8 \times 10^{-5}$ | 1.00 |
| Palmitoleic acid ( $16: 1 \mathrm{n}-7$ ) plasma levels | 11q12.2 | 61,790,331 | C11orf0, C11orf9, FADS1, FADS2, FEN1 | TMEM258 | rs102275-T | $7 \times 10^{-13}$ | $3.8 \times 10^{-5}$ | $3.8 \times 10^{-5}$ | 1.00 |
| Stearic acid (18:0) plasma levels | 11q12.2 | 61,790,331 | C11orf10, FADS1, FADS2, FEN1, FADS3 | TMEM258 | rs102275-C | $1 \times 10^{-20}$ | $3.8 \times 10^{-5}$ | $3.8 \times 10^{-5}$ | 1.00 |
| Crohn's disease | 11q12.2 | 61,790,331 | FADS1 | TMEM258 | rs102275-C | $2 \times 10^{-11}$ | $3.8 \times 10^{-5}$ | $3.8 \times 10^{-5}$ | 1.00 |
| Inflammatory bowel disease | 11q12.2 | 61,796,827 | C11orf9,FADS1,FADS2 | FEN1 | rs4246215-T | $2 \times 10^{-15}$ | $4.0 \times 10^{-6}$ | $4.0 \times 10^{-6}$ | 1.00 |
| Platelet counts | 11q12.2 | 61,796,827 | FEN1 | FEN1 | rs4246215-T | $3 \times 10^{-10}$ | $4.0 \times 10^{-6}$ | $4.0 \times 10^{-6}$ | 1.00 |
| Cholesterol, total | 11 q 12.2 | 61,802,358 | FADS1, FADS2, FADS3 | FADS1 | rs174546-T | $3 \times 10^{-37}$ | $1.8 \times 10^{-5}$ | $1.8 \times 10^{-5}$ | 1.00 |
| HDL cholesterol | 11q12.2 | 61,802,358 | FADS1, FADS2, FADS3 | FADS1 | rs174546-T | $8 \times 10^{-28}$ | $1.8 \times 10^{-5}$ | $1.8 \times 10^{-5}$ | 1.00 |
| LDL cholesterol | 11q12.2 | 61,802,358 | FADS1, FADS2, FADS3 | FADS1 | rs174546-T | $2 \times 10^{-39}$ | $1.8 \times 10^{-5}$ | $1.8 \times 10^{-5}$ | 1.00 |
| Triglycerides | 11q12.2 | 61,802,358 | FADS1, FADS2, FADS3 | FADS1 | rs174546-T | $7 \times 10^{-38}$ | $1.8 \times 10^{-5}$ | $1.8 \times 10^{-5}$ | 1.00 |


| Information from the NHGRI GWAS catalog ${ }^{\text {a }}$ |  |  |  |  |  |  | Information from TAGC asthma meta-analysis |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Disease.Trait | Region | Chr_position (build 38) | Reported Genes | Mapped Gene | Strongest SNP -Risk Allele | P_Value | $P_{\text {fixed }}{ }^{\text {b }}$ | $P_{\text {random }}{ }^{\text {b }}$ | $P_{\text {het }}{ }^{\text {c }}$ |
| Plasma omega-6 polyunsaturated fatty acid levels (arachidonic acid) | 11q12.2 | 61,803,311 | FADS2, C11orf9, FEN1, FADS1, FADS3, RAB3IL1, BEST1, DAGLA, FTH1, INCENP, SYT7, SCGB2A1, SCGB1D1, AHNAK, SCGB2A2 | FADS1 | rs174547-C | OE+00 | $2.1 \times 10^{-5}$ | $2.1 \times 10^{-5}$ | 1.00 |
| Plasma omega-6 polyunsaturated fatty acid levels (gamma-linolenic acid) | 11q12.2 | 61,803,311 | FADS1, DAGLA, C11orf9, C11orf10, FEN1, FADS2, FADS3, RAB3IL1, FTH1, DAGLA, BEST1 | FADS1 | rs174547-T | $2 \times 10^{-72}$ | $2.1 \times 10^{-5}$ | $2.1 \times 10^{-5}$ | 1.00 |
| Metabolic traits | 11q12.2 | 61,803,311 | FADS1 | FADS1 | rs174547-C | $9 \times 10^{-116}$ | $2.1 \times 10^{-5}$ | $2.1 \times 10^{-5}$ | 1.00 |
| Resting heart rate | 11q12.2 | 61,803,311 | FADS1 | FADS1 | rs174547-C | $2 \times 10^{-9}$ | $2.1 \times 10^{-5}$ | $2.1 \times 10^{-5}$ | 1.00 |
| Lipid metabolism phenotypes | 11q12.2 | 61,803,311 | FADS1, FADS2, FADS3 | FADS1 | rs174547-? | $8 \times 10^{-26} 2$ | $2.1 \times 10^{-5}$ | $2.1 \times 10^{-5}$ | 1.00 |
| Heart rate | 11q12.2 | 61,803,910 | FADS1 | FADS1 | rs174549-A | $1 \times 10^{-22}$ | $3.7 \times 10^{-5}$ | $3.7 \times 10^{-5}$ | 1.00 |
| Metabolite levels | 11q12.2 | 61,803,910 | FADS1 | FADS1 | rs174549-G | $2 \times 10^{-30}$ | $3.7 \times 10^{-5}$ | $3.7 \times 10^{-5}$ | 1.00 |
| Fasting glucose-related traits | 11q12.2 | 61,804,006 | FADS1 | FADS1 | rs174550-T | $2 \times 10^{-15}$ | $2.1 \times 10^{-5}$ | $2.1 \times 10^{-5}$ | 1.00 |
| Fasting glucose-related traits (interaction with BMI) | 11q12.2 | 61,804,006 | FADS1 | FADS1 | rs174550-? | $2 \times 10^{-9}$ | $2.1 \times 10^{-5}$ | $2.1 \times 10^{-5}$ | 1.00 |
| Rheumatoid arthritis | 11q12.2 | 61,828,092 | FADS1, FADS2, FADS3 | FADS2 | rs968567-C | $2 \times 10^{-8}$ | $4.2 \times 10^{-4}$ | $4.2 \times 10^{-4}$ | 0.95 |
| P wave duration | 11q12.2 | 61,837,342 | FADS2, C11orf9 | FADS2 | rs174577-A | $3 \times 10^{-8}$ | $2.7 \times 10^{-4}$ | $2.7 \times 10^{-4}$ | 0.91 |
| Liver enzyme levels (alkaline phosphatase) | 11q12.2 | 61,855,668 | C11orf10, FADS1, FADS2 | FADS2 | rs174601-T | $3 \times 10^{-9}$ | $2.6 \times 10^{-4}$ | $2.6 \times 10^{-4}$ | 0.95 |
| Atopic dermatitis | 11q13.1 | 65,784,486 | OVOL1 | AP5B1-OVOL1 | rs479844-G | $1 \times 10^{-13}$ | $3.0 \times 10^{-5}$ | $3.0 \times 10^{-5}$ | 0.48 |
| Atopic dermatitis | 11q13.5 | 76,559,639 | C11orf30,LRRC32 | C11orf30-LRRC32 | rs7130588-G | $4 \times 10^{-13}$ | $2.5 \times 10^{-11}$ | $3.4 \times 10^{-11}$ | 0.46 |
| Allergic sensitization | 11q13.5 | 76,588,150 | C11orf30, LRRC32 | C11orf30-LRRC32 | rs2155219-T | $1 \times 10^{-18}$ | $1.1 \times 10^{-17}$ | $5.1 \times 10^{-13}$ | 0.08 |
| Self-reported allergy | 11q13.5 | 76,588,150 | C11orf30, LRRC32 | C11orf30-LRRC32 | rs2155219-T | $2 \times 10^{-19}$ | $1.1 \times 10^{-17}$ | $5.1 \times 10^{-13}$ | 0.08 |
| Inflammatory bowel disease | 11q13.5 | 76,588,150 | Intergenic | C11orf30-LRRC32 | rs2155219-T | $4 \times 10^{-36}$ | $1.1 \times 10^{-17}$ | $5.1 \times 10^{-13}$ | 0.08 |
| Allergic rhinitis | 11q13.5 | 76,588,150 | C11orf30, LRRC32 | C11orf30-LRRC32 | rs2155219-T | $4 \times 10^{-8}$ | $1.1 \times 10^{-17}$ | $5.1 \times 10^{-13}$ | 0.08 |
| IgE grass sensitization | $11 q 13.5$ | 76,588,150 | C11orf30, LRRC32 | C11orf30-LRRC32 | rs2155219-T | $1 \times 10^{-8}$ | $1.1 \times 10^{-17}$ | $5.1 \times 10^{-13}$ | 0.08 |
| Ulcerative colitis | 11q13.5 | 76,588,150 | Intergenic | C11orf30-LRRC32 | rs2155219-T | $5 \times 10^{-16}$ | $1.1 \times 10^{-17}$ | $5.1 \times 10^{-13}$ | 0.08 |
| Crohn's disease | 11q13.5 | 76,590,331 | C11orf30 | C11orf30-LRRC32 | rs7927997-T | $6 \times 10^{-13}$ | $3.2 \times 10^{-13}$ | $3.2 \times 10^{-13}$ | 0.53 |
| Inflammatory bowel disease | 12q13.11 | 47,814,585 | VDR | HDAC7 | rs11168249-C | $8 \times 10^{-9}$ | $4.4 \times 10^{-5}$ | $5.3 \times 10^{-4}$ | 0.19 |
| Vitiligo | 12q13.2 | 56,007,301 | PMEL, DGKA, CDK2, RAB5B, SUOX, ZNFN1A4, RPS26, ERBB3, PA2G4 | IKZF4 | rs10876864-G | $8 \times 10^{-12}$ | $8.9 \times 10^{-8}$ | $3.2 \times 10^{-4}$ | $3.0 \times 10^{-3}$ |
| Allergic sensitization | 12 q 13.3 | 57,095,926 | STAT6 | NAB2;STAT6 | rs1059513-T | $1 \times 10^{-14}$ | $3.9 \times 10^{-7}$ | $3.9 \times 10^{-7}$ | 0.65 |
| IgE levels | 12 q 13.3 | 57,095,926 | STAT6, NAB2 | NAB2;STAT6 | rs1059513-C | $2 \times 10^{-12}$ | $3.9 \times 10^{-7}$ | $3.9 \times 10^{-7}$ | 0.65 |
| C-reactive protein levels | 12q24.31 | 120,986,153 | HNF1A, C12orf43, OASL | HNF1A | rs2393791-G | $3 \times 10^{-10}$ | $8.6 \times 10^{-5}$ | $8.7 \times 10^{-4}$ | 0.21 |
| C-reactive protein and white blood cell count | 12q24.31 | 120,986,153 | HNF1A | HNF1A | rs2393791-C | $3 \times 10^{-9}$ | $8.6 \times 10^{-5}$ | $8.7 \times 10^{-4}$ | 0.21 |
| Gamma glutamyl transpeptidase | 12q24.31 | 120,986,153 | HNF1A | HNF1A | rs2393791-G | $7 \times 10^{-30}$ | $8.6 \times 10^{-5}$ | $8.7 \times 10^{-4}$ | 0.21 |
| Liver enzyme levels (gamma-glutamyl transferase) | 12q24.31 | 120,987,058 | HNF1A, C12orf27 | HNF1A | rs7310409-G | $7 \times 10^{-45}$ | $8.4 \times 10^{-5}$ | $8.5 \times 10^{-4}$ | 0.21 |
| C-reactive protein | 12q24.31 | 120,987,058 | HNF1A | HNF1A | rs7310409-G | $3 \times 10^{-8}$ | $8.4 \times 10^{-5}$ | $8.5 \times 10^{-4}$ | 0.21 |
| Head circumference (infant) | 12924.31 | 123,338,164 | SBNO1 | SBNO1 | rs7980687-A | $8 \times 10^{-9}$ | $3.2 \times 10^{-5}$ | $8.6 \times 10^{-4}$ | 0.06 |
| Primary biliary cirrhosis | 14 q 24.1 | 68,286,876 | RAD51L1 | RAD51B | rs911263-T | $2 \times 10^{-11}$ | $1.1 \times 10^{-5}$ | $1.1 \times 10^{-5}$ | 0.97 |
| Age-related macular degeneration | 14q24.1 | 68,318,360 | RAD51B | RAD51B | rs8017304-A | $9 \times 10^{-11}$ | $3.9 \times 10^{-5}$ | $3.9 \times 10^{-5}$ | 0.78 |
| Primary tooth development (number of teeth) | 14 q 24.1 | 68,322,207 | RAD51L1 | RAD51B | rs1956529-T | $3 \times 10^{-8}$ | $7.3 \times 10^{-5}$ | $7.3 \times 10^{-5}$ | 0.84 |
| Height | 14q24.1 | 68,346,398 | RAD51L1 | RAD51B | rs1570106-T | $8 \times 10^{-9}$ | $2.0 \times 10^{-4}$ | $2.7 \times 10^{-4}$ | 0.42 |
| Inflammatory bowel disease | 15922.33 | 67,150,258 | SMAD3 | SMAD3 | rs17293632-T | $6 \times 10^{-16}$ | $1.9 \times 10^{-15}$ | $1.5 \times 10^{-11}$ | 0.18 |
| Crohn's disease | 15922.33 | 67,150,258 | SMAD3 | SMAD3 | rs17293632-T | $3 \times 10^{-19}$ | $1.9 \times 10^{-15}$ | $1.5 \times 10^{-11}$ | 0.18 |
| Self-reported allergy | 15q22.33 | 67,157,967 | SMAD3 | SMAD3 | rs17228058-G | $1 \times 10^{-8}$ | $5.4 \times 10^{-15}$ | $9.4 \times 10^{-11}$ | 0.14 |
| Corneal structure | 15922.33 | 67,175,169 | SMAD3 | SMAD3 | rs12913547-T | $5 \times 10^{-10}$ | $9.1 \times 10^{-6}$ | $3.5 \times 10^{-5}$ | 0.27 |
| Height | 15925.2 | 83,911,404 | ADAMTSL3 | ADAMTSL3 | rs11259933-A | $1 \times 10^{-19}$ | $7.3 \times 10^{-5}$ | $7.3 \times 10^{-5}$ | 0.83 |
| Type 1 diabetes autoantibodies | 16p13.13 | 11,086,016 | CLEC16A | CLEC16A | rs12708716-G | $5 \times 10^{-14}$ | $1.1 \times 10^{-7}$ | $4.5 \times 10^{-5}$ | 0.07 |
| Primary biliary cirrhosis | 16p13.13 | 11,093,926 | CLEC16A | CLEC16A | rs12924729-G | $3 \times 10^{-12}$ | $4.2 \times 10^{-6}$ | $4.0 \times 10^{-4}$ | 0.07 |
| Type 1 diabetes | 16 p 13.13 | 11,144,926 | KIAA0350 | CLEC16A | rs2903692-G | $7 \times 10^{-11}$ | $2.4 \times 10^{-7}$ | $1.8 \times 10^{-4}$ | 0.04 |
| Rheumatoid arthritis | 17q12 | 39,583,908 | MED1 | CDK12-NEUROD2 | rs1877030-C | $2 \times 10^{-8}$ | $2.6 \times 10^{-13}$ | $2.6 \times 10^{-13}$ | 0.99 |
| Primary biliary cirrhosis | 17q12 | 39,820,216 | IKZF3, ZPBP2, GSDMB, ORMDL3 | IKZF3 | rs9303277-T | $4 \times 10^{-9}$ | $4.9 \times 10^{-43}$ | $9.6 \times 10^{-26}$ | 0.02 |
| Ulcerative colitis | 17q12 | 39,884,510 | IKZF3, ORMDL3, IKZF3, PNMT, ZPBP2, GSDML | ZPBP2-GSDMB | rs2872507-A | $5 \times 10^{-11}$ | $8.1 \times 10^{-38}$ | $2.0 \times 10^{-24}$ | 0.04 |
| Crohn's disease | 17q12 | 39,884,510 | GSMDL,ZPBP2,ORMDL3,IKZF3 | ZPBP2-GSDMB | rs2872507-A | $2 \times 10^{-9}$ | $8.1 \times 10^{-38}$ | $2.0 \times 10^{-24}$ | 0.04 |
| Cervical cancer | 17q12 | 39,895,095 | GSDMB | ZPBP2-GSDMB | rs8067378-G | $9 \times 10^{-10}$ | $4.8 \times 10^{-42}$ | $1.5 \times 10^{-28}$ | 0.07 |
| Fractional exhaled nitric oxide (childhood) | 17 q 12 | 39,900,944 | ZPBP2, GSDMB | ZPBP2-GSDMB | rs8069176-A | $2 \times 10^{-8}$ | $4.6 \times 10^{-41}$ | $2.0 \times 10^{-23}$ | $6.1 \times 10^{-3}$ |
| Type 1 diabetes | 17q12 | 39,909,987 | ORMDL3 | GSDMB | rs2290400-? | $6 \times 10^{-13}$ | $1.5 \times 10^{-43}$ | $1.5 \times 10^{-25}$ | 0.01 |


| Information from the NHGRI GWAS catalog ${ }^{\text {a }}$ |  |  |  |  |  |  | Information from TAGC asthma meta-analysis |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Disease.Trait | Region | Chr_position (build 38) | Reported Genes | Mapped Gene | Strongest SNP -Risk Allele | P_Value | $P_{\text {fixed }}{ }^{\text {b }}$ | $P_{\text {random }}{ }^{\text {b }}$ | $\boldsymbol{P}_{\text {het }}{ }^{\text {c }}$ |
| Self-reported allergy | 17 q 12 | 39,917,778 | GSDMB, IKZF3 | GSDMB | rs9303280-T | $9 \times 10^{-9}$ | $3.0 \times 10^{-43}$ | $9.3 \times 10^{-25}$ | 0.01 |
| Hematological parameters | 17q21.1 | 39,954,436 | GSDMA, ORMDL3 | LRRC3C-GSDMA | rs17609240-G | $9 \times 10^{-9}$ | $2.3 \times 10^{-23}$ | $5.8 \times 10^{-12}$ | 0.03 |
| White blood cell types | 17q21.1 | 40,000,459 | PSMD3, CSF3 | PSMD3-CSF3 | rs4794822-T | $4 \times 10^{-16}$ | $2.0 \times 10^{-16}$ | $1.9 \times 10^{-11}$ | 0.09 |
| Neutrophil count | 17q21.1 | 40,000,459 | PSMD3, CSF3 | PSMD3-CSF3 | rs $4794822-\mathrm{C}$ | $6 \times 10^{-10}$ | $2.0 \times 10^{-16}$ | $1.9 \times 10^{-11}$ | 0.09 |
| C-reactive protein and white blood cell count | 17 q 21.1 | 40,010,626 | Intergenic | PSMD3-CSF3 | rs8078723-C | $3 \times 10^{-9}$ | $9.1 \times 10^{-17}$ | $1.6 \times 10^{-11}$ | 0.08 |
| White blood cell count | 17q21.1 | 40,010,626 | GSDMB, ORMDL3, GSDMA, PSMD3, CSF3, MED24, SNORD124, THRA, NR1D1 | PSMD3-CSF3 | rs8078723-T | $2 \times 10^{-31}$ | $9.1 \times 10^{-17}$ | $1.6 \times 10^{-11}$ | 0.08 |
| Diastolic blood pressure | 17q21.33 | 49,363,104 | ZNF652, PHB | ZNF652;LOC102724596 | rs16948048-G | $5 \times 10^{-9}$ | $7.2 \times 10^{-9}$ | $1.9 \times 10^{-8}$ | 0.39 |
| Sex hormone-binding globulin levels | 17q21.33 | 49,368,389 | ZNF652 | LOC102724596 | rs2411984-A | $4 \times 10^{-14}$ | $5.6 \times 10^{-5}$ | $8.4 \times 10^{-4}$ | 0.10 |

ariables (highlighted in blue) are defined in the catalog web site: https://www. obiac.uk/gwas/
${ }^{\text {a }}$ Variables extracted from the GWAS catalog (highlighted in blue) are defined in the catalog web site: https://www.ebi.ac.uk/gwas/
${ }^{\text {b }}$ TAGC Variables are highlighted in orange: $\mathrm{P}_{\text {fixed }}$ and $\mathrm{P}_{\text {random }}$ are the P -values for association with asthma in multi-ancestry meta-analysis under fixed-effects and random-effects models, respectively
${ }^{c} P_{\text {het }}$ is the $P$-value for heterogeneity in SNP effect size across TAGC studies using the Cochran's $Q$ test

## Supplementary Table 17. Enrichment of asthma risk SNPs in promoter and enhancer marks by cell type

The results presented in this table are for 16 out of the 18 asthma loci shown in Table 1
The 6 p21.33 and 6 p21.32 loci spanning the HLA complex were excluded because of high variability and LD in the region
The method used to assess enrichment is described in detail in the Online Methods
The enhancer and promoter histone marks were defined using a 15 state model applied to ROADMAP and ENCODE data (PMID:25693563; http://egg2.wustl.edu/roadmap/data)


|  |  |  |  |  | Active promoters |  | Active and inactive promoters |  | Active enhancers |  | Active and inactive enhancers |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| EID ${ }^{\text {a }}$ | Group ${ }^{\text {a }}$ | Standardized Epigenome name ${ }^{\text {a }}$ | Anatomy ${ }^{\text {a }}$ | Type ${ }^{\text {a }}$ | $\begin{aligned} & \text { AST } \\ & \text { locib }^{\text {ach }} \end{aligned}$ | FDR ${ }^{\text {c }}$ | $\begin{aligned} & \text { AST } \\ & \text { loci }^{\text {b }} \end{aligned}$ | FDR ${ }^{\text {c }}$ | $\begin{aligned} & \text { AST } \\ & \text { locib } \end{aligned}$ | FDR ${ }^{\text {c }}$ | $\begin{aligned} & \text { AST } \\ & \text { locib }^{2} \end{aligned}$ | FDR ${ }^{\text {c }}$ |
| E040 | Blood \& T-cell | Primary T helper memory cells from peripheral blood 1 | BLOOD | PrimaryCell | 6 | 0.079 | 6 | 0.114 | 11 | 0.001 | 11 | 0.001 |
| E041 | Blood \& T-cell | Primary T helper cells PMA-I stimulated | BLOOD | PrimaryCell | 7 | 0.070 | 7 | 0.098 | 10 | 0.001 | 10 | 0.004 |
| E042 | Blood \& T-cell | Primary $T$ helper 17 cells PMA-I stimulated | BLOOD | PrimaryCell | 4 | 0.224 | 4 | 0.260 | 10 | 0.001 | 10 | 0.002 |
| E043 | Blood \& T-cell | Primary $T$ helper cells from peripheral blood | BLOOD | PrimaryCell | 5 | 0.191 | 5 | 0.219 | 11 | 0.001 | 11 | 0.002 |
| E044 | Blood \& T-cell | Primary T regulatory cells from peripheral blood | BLOOD | PrimaryCell | 7 | 0.060 | 7 | 0.074 | 11 | 0.001 | 11 | 0.001 |
| E045 | Blood \& T-cell | Primary T cells effector/memory enriched from peripheral blood | BLOOD | PrimaryCell | 7 | 0.060 | 7 | 0.074 | 11 | 0.001 | 11 | 0.001 |
| E046 | HSC \& B-cell | Primary Natural Killer cells from peripheral blood | BLOOD | PrimaryCell | 6 | 0.075 | 6 | 0.114 | 9 | 0.010 | 9 | 0.017 |
| E047 | Blood \& T-cell | Primary T CD8+ naive cells from peripheral blood | BLOOD | PrimaryCell | 5 | 0.171 | 5 | 0.203 | 10 | 0.001 | 10 | 0.003 |
| E048 | Blood \& T-cell | Primary T CD8+ memory cells from peripheral blood | BLOOD | PrimaryCell | 7 | 0.060 | 7 | 0.074 | 11 | 0.001 | 11 | 0.001 |
| E049 | Mesench | Mesenchymal Stem Cell Derived Chondrocyte Cultured Cells | STROMAL_CONNECTIVE | PrimaryCulture | 5 | 0.289 | 5 | 0.310 | 9 | 0.069 | 11 | 0.013 |
| E050 | HSC \& B-cell | Primary hematopoietic stem cells G-CSF-mobilized Female | BLOOD | PrimaryCell | 6 | 0.144 | 6 | 0.150 | 9 | 0.040 | 9 | 0.059 |
| E051 | HSC \& B-cell | Primary hematopoietic stem cells G-CSF-mobilized Male | BLOOD | PrimaryCell | 7 | 0.127 | 7 | 0.142 | 9 | 0.040 | 9 | 0.058 |
| E052 | Myosat | Muscle Satellite Cultured Cells | MUSCLE | PrimaryCulture | 6 | 0.171 | 6 | 0.208 | 10 | 0.021 | 11 | 0.007 |
| E053 | Neurosph | Cortex derived primary cultured neurospheres | BRAIN | PrimaryCulture | 4 | 0.280 | 4 | 0.290 | 7 | 0.213 | 7 | 0.259 |
| E054 | Neurosph | Ganglion Eminence derived primary cultured neurospheres | BRAIN | PrimaryCulture | 5 | 0.218 | 5 | 0.251 | 6 | 0.222 | 6 | 0.246 |
| E055 | Epithelial | Foreskin Fibroblast Primary Cells skin01 | SKIN | PrimaryCulture | 5 | 0.218 | 6 | 0.208 | 8 | 0.083 | 8 | 0.130 |
| E056 | Epithelial | Foreskin Fibroblast Primary Cells skin02 | SKIN | PrimaryCulture | 4 | 0.364 | 5 | 0.246 | 10 | 0.001 | 10 | 0.005 |
| E057 | Epithelial | Foreskin Keratinocyte Primary Cells skin02 | SKIN | PrimaryCulture | 4 | 0.364 | 5 | 0.250 | 11 | 0.006 | 11 | 0.007 |
| E058 | Epithelial | Foreskin Keratinocyte Primary Cells skin03 | SKIN | PrimaryCulture | 5 | 0.218 | 5 | 0.246 | 11 | 0.006 | 11 | 0.009 |
| E059 | Epithelial | Foreskin Melanocyte Primary Cells skin01 | SKIN | PrimaryCulture | 4 | 0.280 | 4 | 0.305 | 7 | 0.049 | 7 | 0.077 |
| E061 | Epithelial | Foreskin Melanocyte Primary Cells skin03 | SKIN | PrimaryCulture | 3 | 0.468 | 4 | 0.287 | 8 | 0.133 | 8 | 0.166 |
| E062 | Blood \& T-cell | Primary mononuclear cells from peripheral blood | BLOOD | PrimaryCell | 3 | 0.414 | 3 | 0.469 | 8 | 0.001 | 8 | 0.001 |
| E063 | Adipose | Adipose Nuclei | FAT | PrimaryTissue | 5 | 0.224 | 5 | 0.260 | 7 | 0.203 | 7 | 0.241 |
| E065 | Heart | Aorta | VASCULAR | PrimaryTissue | 5 | 0.179 | 5 | 0.208 | 5 | 0.072 | 5 | 0.077 |
| E066 | Other | Liver | LIVER | PrimaryTissue | 5 | 0.218 | 5 | 0.246 | 10 | 0.015 | 10 | 0.016 |
| E067 | Brain | Brain Angular Gyrus | BRAIN | PrimaryTissue | 5 | 0.218 | 5 | 0.256 | 6 | 0.133 | 6 | 0.155 |
| E068 | Brain | Brain Anterior Caudate | BRAIN | PrimaryTissue | 4 | 0.488 | 4 | 0.511 | 8 | 0.040 | 8 | 0.046 |
| E069 | Brain | Brain Cingulate Gyrus | BRAIN | PrimaryTissue | 5 | 0.218 | 5 | 0.260 | 6 | 0.168 | 6 | 0.187 |
| E070 | Brain | Brain Germinal Matrix | BRAIN | PrimaryTissue | 4 | 0.284 | 4 | 0.370 | 5 | 0.269 | 5 | 0.313 |
| E071 | Brain | Brain Hippocampus Middle | BRAIN | PrimaryTissue | 4 | 0.441 | 4 | 0.489 | 6 | 0.238 | 6 | 0.262 |
| E072 | Brain | Brain Inferior Temporal Lobe | BRAIN | PrimaryTissue | 4 | 0.437 | 4 | 0.473 | 5 | 0.301 | 5 | 0.319 |
| E073 | Brain | Brain_Dorsolateral_Prefrontal_Cortex | BRAIN | PrimaryTissue | 5 | 0.218 | 5 | 0.260 | 6 | 0.095 | 6 | 0.110 |
| E074 | Brain | Brain Substantia Nigra | BRAIN | PrimaryTissue | 4 | 0.395 | 4 | 0.420 | 5 | 0.354 | 5 | 0.379 |
| E075 | Digestive | Colonic Mucosa | GI_COLON | PrimaryTissue | 5 | 0.201 | 5 | 0.239 | 6 | 0.023 | 6 | 0.030 |
| E076 | Sm. Muscle | Colon Smooth Muscle | GI_COLON | PrimaryTissue | 5 | 0.213 | 5 | 0.246 | 7 | 0.116 | 7 | 0.133 |
| E077 | Digestive | Duodenum Mucosa | GI_DUODENUM | PrimaryTissue | 7 | 0.106 | 7 | 0.130 | 5 | 0.321 | 6 | 0.187 |
| E078 | Sm. Muscle | Duodenum Smooth Muscle | GI_DUODENUM | PrimaryTissue | 5 | 0.218 | 5 | 0.260 | 6 | 0.099 | 6 | 0.125 |
| E079 | Digestive | Esophagus | GI_ESOPHAGUS | PrimaryTissue | 5 | 0.171 | 5 | 0.208 | 6 | 0.094 | 6 | 0.115 |
| E080 | Other | Fetal Adrenal Gland | ADRENAL | PrimaryTissue | 4 | 0.218 | 4 | 0.261 | 9 | 0.058 | 9 | 0.107 |
| E081 | Brain | Fetal Brain Male | BRAIN | PrimaryTissue | 3 | 0.395 | 3 | 0.421 | 6 | 0.247 | 6 | 0.340 |
| E082 | Brain | Fetal Brain Female | BRAIN | PrimaryTissue | 4 | 0.362 | 4 | 0.426 | 4 | 0.347 | 4 | 0.376 |
| E083 | Heart | Fetal Heart | HEART | PrimaryTissue | 3 | 0.437 | 3 | 0.453 | 8 | 0.167 | 8 | 0.187 |
| E084 | Digestive | Fetal Intestine Large | GI_INTESTINE | PrimaryTissue | 3 | 0.488 | 4 | 0.346 | 7 | 0.133 | 7 | 0.166 |
| E085 | Digestive | Fetal Intestine Small | GI_INTESTINE | PrimaryTissue | 5 | 0.179 | 5 | 0.222 | 5 | 0.495 | 5 | 0.544 |
| E086 | Other | Fetal Kidney | KIDNEY | PrimaryTissue | 5 | 0.193 | 5 | 0.227 | 7 | 0.040 | 7 | 0.049 |
| E087 | Other | Pancreatic Islets | PANCREAS | PrimaryTissue | 4 | 0.426 | 4 | 0.453 | 3 | 0.496 | 3 | 0.501 |
| E088 | Other | Fetal Lung | LUNG | PrimaryTissue | 6 | 0.112 | 6 | 0.136 | 8 | 0.129 | 8 | 0.161 |
| E089 | Muscle | Fetal Muscle Trunk | MUSCLE | PrimaryTissue | 4 | 0.224 | 4 | 0.261 | 6 | 0.123 | 7 | 0.140 |



The enhancer and promoter histone marks were defined using a 15 state model applied to ROADMAP and ENCODE data (PMID:25693563; http://egg2.wustl.edu/roadmap/data)
${ }^{\text {a }}$ EID is the cell type number; the following variables are group, standardized epigenome name, anatomy and type for each cell type (see PMID:25693563 for more details)
${ }^{\text {b }}$ AST_loci is the number of asthma loci that co-localize to a promoter or enhancer mark
${ }^{\text {c }}$ FDR is the Benjamini-Hochberg false discovery rate

## Supplementary Table 18. Enrichment of TAGC asthma-associated loci in DNAse I hypersensitive sites (DHSs) by cell type

The results presented in this table are for 16 out of the 18 asthma loci shown in Table 1
The 6 p21.33 and $6 p 21.32$ loci spanning the HLA complex were excluded because of high variability and LD in the region
The method used to assess enrichment is described in detail in the Online Methods
The DHSs were available in 51 cell types using ROADMAP and ENCODE data (PMID:25693563; http://egg2.wustl.edu/roadmap/data)

| EID ${ }^{\text {a }}$ | Group ${ }^{\text {a }}$ | Standardized Epigenome name ${ }^{\text {a }}$ | Anatomy ${ }^{\text {a }}$ | Type ${ }^{\text {a }}$ | AST_loci ${ }^{\text {b }}$ | FDR ${ }^{\text {c }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| E005 | ES-deriv | H1 BMP4 Derived Trophoblast Cultured Cells | ESC_DERIVED | ESCDerived | 4 | 0.678 |
| E006 | ES-deriv | H1 Derived Mesenchymal Stem Cells | ESC_DERIVED | ESCDerived | 5 | 0.678 |
| E007 | ES-deriv | H1 Derived Neuronal Progenitor Cultured Cells | ESC_DERIVED | ESCDerived | 6 | 0.429 |
| E008 | ESC | H9 Cells | ESC | PrimaryCulture | 10 | 0.024 |
| E017 | IMR90 | IMR90 fetal lung fibroblasts Cell Line | LUNG | CellLine | 3 | 0.920 |
| E021 | iPSC | iPS DF 6.9 Cells | IPSC | PrimaryCulture | 8 | 0.251 |
| E022 | iPSC | iPS DF 19.11 Cells | IPSC | PrimaryCulture | 5 | 0.747 |
| E028 | Epithelial | Breast variant Human Mammary Epithelial Cells (vHMEC) | BREAST | PrimaryCulture | 8 | 0.168 |
| E029 | HSC \& B-cell | Primary monocytes from peripheral blood | BLOOD | PrimaryCell | 3 | 0.744 |
| E032 | HSC \& B-cell | Primary B cells from peripheral blood | BLOOD | PrimaryCell | 8 | 0.057 |
| E033 | Blood \& T-cell | Primary T cells from cord blood | BLOOD | PrimaryCell | 8 | 0.043 |
| E034 | Blood \& T-cell | Primary T cells from peripheral blood | BLOOD | PrimaryCell | 10 | 0.005 |
| E046 | HSC \& B-cell | Primary Natural Killer cells from peripheral blood | BLOOD | PrimaryCell | 8 | 0.048 |
| E050 | HSC \& B-cell | Primary hematopoietic stem cells G-CSF-mobilized Female | BLOOD | PrimaryCell | 7 | 0.232 |
| E051 | HSC \& B-cell | Primary hematopoietic stem cells G-CSF-mobilized Male | BLOOD | PrimaryCell | 0 | 1.000 |
| E055 | Epithelial | Foreskin Fibroblast Primary Cells skin01 | SKIN | PrimaryCulture | 0 | 1.000 |
| E056 | Epithelial | Foreskin Fibroblast Primary Cells skin02 | SKIN | PrimaryCulture | 0 | 1.000 |
| E057 | Epithelial | Foreskin Keratinocyte Primary Cells skin02 | SKIN | PrimaryCulture | 9 | 0.013 |
| E059 | Epithelial | Foreskin Melanocyte Primary Cells skin01 | SKIN | PrimaryCulture | 9 | 0.156 |
| E080 | Other | Fetal Adrenal Gland | ADRENAL | PrimaryTissue | 6 | 0.445 |
| E081 | Brain | Fetal Brain Male | BRAIN | PrimaryTissue | 7 | 0.393 |
| E082 | Brain | Fetal Brain Female | BRAIN | PrimaryTissue | 7 | 0.381 |
| E083 | Heart | Fetal Heart | HEART | PrimaryTissue | 9 | 0.156 |
| E084 | Digestive | Fetal Intestine Large | GI_INTESTINE | PrimaryTissue | 4 | 0.710 |
| E085 | Digestive | Fetal Intestine Small | GI_INTESTINE | PrimaryTissue | 4 | 0.568 |
| E086 | Other | Fetal Kidney | KIDNEY | PrimaryTissue | 6 | 0.523 |
| E088 | Other | Fetal Lung | LUNG | PrimaryTissue | 8 | 0.381 |
| E089 | Muscle | Fetal Muscle Trunk | MUSCLE | PrimaryTissue | 8 | 0.240 |
| E090 | Muscle | Fetal Muscle Leg | MUSCLE_LEG | PrimaryTissue | 9 | 0.161 |
| E091 | Other | Placenta | PLACENTA | PrimaryTissue | 6 | 0.558 |
| E092 | Digestive | Fetal Stomach | GI_STOMACH | PrimaryTissue | 6 | 0.438 |
| E093 | Thymus | Fetal Thymus | THYMUS | PrimaryTissue | 8 | 0.130 |
| E094 | Digestive | Gastric | GI_STOMACH | PrimaryTissue | 3 | 0.710 |
| E097 | Other | Ovary | OVARY | PrimaryTissue | 5 | 0.541 |
| E098 | Other | Pancreas | PANCREAS | PrimaryTissue | 5 | 0.541 |
| E100 | Muscle | Psoas Muscle | MUSCLE | PrimaryTissue | 5 | 0.592 |
| E109 | Digestive | Small Intestine | GI_INTESTINE | PrimaryTissue | 6 | 0.393 |
| E114 | ENCODE2012 | A549 EtOH 0.02pct Lung Carcinoma Cell Line | LUNG | Celline | 3 | 0.839 |
| E116 | ENCODE2012 | GM12878 Lymphoblastoid Cells | BLOOD | PrimaryCulture | 9 | 0.043 |
| E117 | ENCODE2012 | HeLa-S3 Cervical Carcinoma Cell Line | CERVIX | CellLine | 8 | 0.148 |
| E118 | ENCODE2012 | HepG2 Hepatocellular Carcinoma Cell Line | LIVER | Cellline | 4 | 0.678 |
| E119 | ENCODE2012 | HMEC Mammary Epithelial Primary Cells | BREAST | PrimaryCulture | 9 | 0.076 |
| E120 | ENCODE2012 | HSMM Skeletal Muscle Myoblasts Cells | MUSCLE | PrimaryCulture | 6 | 0.541 |
| E121 | ENCODE2012 | HSMM cell derived Skeletal Muscle Myotubes Cells | MUSCLE | PrimaryCulture | 7 | 0.449 |
| E122 | ENCODE2012 | HUVEC Umbilical Vein Endothelial Primary Cells | VASCULAR | PrimaryCulture | 7 | 0.161 |
| E123 | ENCODE2012 | K562 Leukemia Cells | BLOOD | PrimaryCulture | 7 | 0.240 |
| E124 | ENCODE2012 | Monocytes-CD14+ R001746 Primary Cells | BLOOD | PrimaryCell | 6 | 0.322 |
| E125 | ENCODE2012 | NH-A Astrocytes Primary Cells | BRAIN | PrimaryCulture | 8 | 0.168 |
| E126 | ENCODE2012 | NHDF-Ad Adult Dermal Fibroblast Primary Cells | SKIN | PrimaryCulture | 5 | 0.710 |
| E127 | ENCODE2012 | NHEK-Epidermal Keratinocyte Primary Cells | SKIN | PrimaryCulture | 7 | 0.245 |
| E128 | ENCODE2012 | NHLF Lung Fibroblast Primary Cells | LUNG | PrimaryCulture | 6 | 0.496 |

${ }^{\text {a }}$ EID is the cell type number; the following variables are group, standardized epigenome name, anatomy and type for each cell type (PMID:25693563)
${ }^{\mathrm{b}}$ AST_loci is the number of asthma loci that co-localize to a DHSs
${ }^{\text {c F FDR }}$ is the Benjamini-Hochberg false discovery rate

Supplementary Table 19. Results of GRAIL analysis for genome-wide significant asthma-associated loci The 18 TAGC asthma-associated loci are ordered according to $P_{\text {GRAIL }}$ from lowest to highest, see Online Methods for details on GRAIL method (PMID: 19557189); no gene was found by GRAIL at 10p14 locus

| Region | Lead SNP | Start ${ }^{\text {a }}$ | Stop ${ }^{\text {a }}$ | $\begin{gathered} \text { PGrail }^{\text {b }} \\ \text { for the region } \end{gathered}$ |
| :---: | :---: | :---: | :---: | :---: |
| 5q22.1 | rs10455025 | 110,429,500 | 110,519,500 | 8.65E-08 |
| 9 p 24.1 | rs992969 | 6,134,500 | 6,207,000 | $5.39 \mathrm{E}-07$ |
| 12q13.3 | rs167769 | 55,606,500 | 55,835,500 | $1.63 \mathrm{E}-06$ |
| 2 q 12 | rs1420101 | 102,282,914 | 102,461,414 | $1.75 \mathrm{E}-06$ |
| 5q31 | rs20541 | 131,904,000 | 132,034,500 | 3.84E-06 |
| 6p21.32 | rs9272346 | 32,447,500 | 32,790,500 | $2.64 \mathrm{E}-04$ |
| 16p13.13 | rs17806299 | 10,925,500 | 11,215,000 | 5.82E-04 |
| 6 p 21.33 | rs3131064 | 30,332,000 | 31,536,500 | $1.07 \mathrm{E}-03$ |
| 6p22.1 | rs1233578 | 28,283,500 | 29,463,500 | $1.24 \mathrm{E}-03$ |
| 17q12-q21 | rs2952156 | 34,634,000 | 35,343,500 | $1.28 \mathrm{E}-02$ |
| $11 q 13.5$ | rs7927894 | 75,802,500 | 76,016,000 | $1.81 \mathrm{E}-02$ |
| $6 q 15$ | rs2325291 | 90,867,000 | 91,080,500 | $5.01 \mathrm{E}-02$ |
| $17 q 21.33$ | rs17637472 | 44,675,500 | 44,834,000 | $5.20 \mathrm{E}-02$ |
| 8 q 21.13 | rs12543811 | 81,052,000 | 81,745,500 | $8.25 \mathrm{E}-02$ |
| $15 q 22.2$ | rs11071558 | 58,803,500 | 58,922,000 | $5.80 \mathrm{E}-01$ |
| 5q31.3 | rs7705042 | 141,394,500 | 141,546,000 | $6.14 \mathrm{E}-01$ |
| 15q22.33 | rs2033784 | 65,225,500 | 65,255,500 | $6.54 \mathrm{E}-01$ |
| 10p14 | rs2589561 | 8,828,500 | 9,406,000 | NA |

aStart and stop are the start and stop positions of the regions defined by GRAIL ${ }^{\mathrm{b}} P_{\text {GRAIL }}$ measures the strength of relatedness of each region with the other regions estimated by mining PubMed abstracts; the P-value is corrected for the number of genes assigned to that region by GRAIL.

## Supplementary Note

Study descriptions, Funding and acknowledgements, and Contributing groups

## Study Descriptions

All studies obtained approval from their local ethical committees. All participants provided written informed consent. The ethical statement will not be provided for each individual study

## Australian Asthma Genetics Consortium (AAGC)

Subjects: Participants were drawn from two cohorts described in greater detail elsewhere ${ }^{1}$ : the Australian Asthma Genetics Consortium (AAGC) cohort ( $n=1,810$ ) and the Queensland Institute of Medical Research (QIMR) GWAS cohort ( $n=4,157$ ). Of the 2110 asthmatic patients, 1,269 ( $60 \%$ ) were classified as having childhood asthma (defined by an age-of-onset at or before age 16 years) 515 (24\%) patients with later onset asthma (age-of-onset after the age of 16 years) and 326 (16\%) with unknown age-of-onset. 1,232 (58\%) of asthmatics were atopic, as defined by a positive skin prick test response to at least one common allergen; 1,444 (68\%) had at least one first-degree relative with asthma; and 760 (36\%) reported lifetime smoking. The 3,857 controls included $2,030(53 \%)$ individuals who were classified as asthma-free based on clinical examination (109 [3\%]) or epidemiological questionnaires (2,592 [50\%]). The remaining 1,827 (47\%) individuals provided no information about their asthma status but were included in the analysis to improve power. Mean age was 34 years (SD 16.2, range 2-92) and 3,267 (55\%) were women. All participants were confirmed to be unrelated and of European ancestry based on genetic data.

Genotyping and QC: The Australian GWAS included data for 4,293 (72\%) individuals genotyped with Illumina 610K array and 1,674 (28\%) with Illumina 370K array ${ }^{1}$. The same single-nucleotide polymorphism (SNP) quality control filters were applied to each dataset individually, including the removal of SNPs with call rate lower than 95\%, minor allele frequency (MAF) lower than 0.01, and Hardy-Weinberg equilibrium test $\mathrm{p}<10^{-6}$.

Data Imputation and Analysis: Autosomal SNPs passing quality control were used to impute $\mathbf{7 . 8}$ million variants available from the combined 1000 genomes ( 60 individuals from the CEU collection; March 2010 release) and HapMap 3 ( 955 individuals from 11 populations; February 2009 release) reference panels with IMPUTE2 ${ }^{2}$. The 610K and 370K datasets were imputed separately. After imputation, we excluded SNPs with imputation information $<0 \cdot 3$, MAF $<0 \cdot 01$, or Hardy-Weinberg equilibrium test $p<10^{-6}$. We also excluded from analysis SNPs with significant ( $p<0 \cdot 001$ ) allele frequency differences between the two imputation analysis groups (from case-case and control-control comparisons). After quality control, genotype data for 5.7 million SNPs were merged across the 610 K and 370 K datasets; only results for Hapmap2 SNPS were provided to TAGC. Individual SNPs were then tested for association with lifetime
physician-diagnosed asthma with a Cochran-Mantel-Haenszel test as implemented in PLINK ${ }^{3}$, with two strata corresponding to the two imputation analysis groups.

## ALLERGEN Consortium

Canadian Cohorts (see http://www.genopha.ca/)
Subjects: The Canadian Asthma Primary Prevention Study (CAPPS ${ }^{4}$ ) was initiated in 1995 to assess the effectiveness of a multifaceted intervention program on the primary prevention of asthma in high-risk infants. Infants at high risk for developing asthma and other allergic disorders were identified and their mothers were recruited during their third trimester of pregnancy. Infants at high risk were defined as those who had a parent with asthma or two first-degree relatives with allergies or atopic disorders. The study had two recruitment centers in Canada, (Vancouver and Winnipeg). In total, there were 545 families recruited into the study ( 549 infants, 4 sets of twins). Loss to follow-up was minimal, with $86 \%$ of the families completing the questionnaire at the 7 -year time point. There have been numerous publications on this cohort evaluating the effectiveness of the intervention strategy. For genetic analysis blood samples have been obtained from the children, and both parents.

The Study of Asthma Genes and Environment Study (SAGE ${ }^{5}$ ) is a population-based sample of 16,320 children, born in the province of Manitoba, Canada in the year of 1995. In 2002, a one-page survey was mailed to families to enquire about their health and home environment exposure. Questions included several chronic conditions in childhood, including hayfever, food allergy and asthma. Parents were asked for permission to be contacted for further study and to link the survey data to their child's health care database records. Children were stratified according to the presence of asthma ( $n=392$ ), to the presence of hayfever or food allergy ( $n=192$ ) and neither ( $n=3,002$ ). Children with neither condition were further grouped by rural and urban location; the latter were further stratified by low and high income neighborhoods. All children in the asthma and allergy strata were invited to participate in the case-control study, as were a random sample $(\mathrm{n}=200)$ from the strata of children with neither condition. Following consent from parents to participate in the study, children were assessed for allergic phenotypes.

After QC a total of 766 subjects from CAPPS were retained analyses ( 360 cases/ 406 controls) and a total of 1022 subjects from SAGE (353 cases / 669 controls).

Genotyping and QC: Genome-wide SNP genotyping was performed using the Illumina Human610 quad array of which 821 subjects from the Canadian Cohorts (CAPPS $n=414$ and SAGE $n=407$ ) were included in the GABRIEL consortium GWAS analyses ${ }^{6}$, using the GABRIEL QC protocol. The remaining 967 subjects (CAPPS $n=352$ and SAGE $n=615$ ) were genotyped using the same chip (Illumina Human610 quad array) and
the following QC protocols: < $97 \%$ call rate for both samples and SNPs, evaluation of Hardy-Weinberg Equilibrium (HWE), heterozygosity (3 s.d.), ethnicity checks using multidimensional scaling, checks for duplicate samples, verification of reported family relationships and Mendelian errors.
Data Imputation and Analysis: Imputation was performed using MACH with HapMap2 CEU as the reference population. Statistical analyses were performed using STATA (childhood/offspring generation) to account for sibling relationships and PLINK (case-control) for adults/parents.

## Saguenay Lac Saint Jean Study (SLSJ)

Subjects: The Saguenay-Lac-Saint-Jean (SLSJ) asthma study, consists of French-Canadian families ascertained through asthmatic probands ${ }^{7}$. Probands were included in the study if they fulfill at least two of the following criteria: 1) a minimum of three clinic visits for acute asthma within one year; 2) two or more asthma-related hospital admissions within one year; or 3) steroid dependency, as defined by either six months of oral, or one year of inhaled corticosteroid use. Individuals from 253 independent families were included in the study. Families were included in the study if at least one parent was available for phenotypic assessment, at least one parent was unaffected, and all four grandparents were of FrenchCanadian origin. Family members were considered asthmatics if both a self-reported history of asthma and a history of physician-diagnosed asthma were available, or by clinical evaluation following a methacholine provocation test. After QC a total of 1,199 subjects were retained in the GWAS analyses.
Genotyping and QC: Genome-wide SNP genotyping was performed using the Illumina Human610 Quad array, as part of the GABRIEL consortium (see GABRIEL paragraph below for details).

Data Imputation and Analysis: Imputation and analysis were done as part of the GABRIEL consortium (see GABRIEL paragraph below).

## APCAT Consortium

The APCAT Consortium includes 21,858 European-ancestry subjects ( 2,055 physician-diagnosed asthmatics and 19,803 asthma-free controls) with available genome-wide data that participated in seven population based-studies, five studies from Finland (the Northern Finland Birth Cohort 1966 [NFBC66]; Helsinki Birth Cohort [HBC]; Health 2000 [H2000]; FINRISK, including FINRISK 1992, 1997, 2002 and 2007; and the Young Finns Study), the European Prospective Investigation of Cancer, Norfolk (EPIC-Norfolk) and Framingham Heart Study from the United States.

## Framingham Heart Study:

Subjects: The FHS is a collection of cohorts recruited to investigate cardiovascular disease and its risk factors as described in detail elsewhere ${ }^{8}$. Asthma was classified based on self-report of a physician's diagnosis; 797 asthmatics and 6,463 controls were included in TAGC study.

Genotyping and QC: Genome-wide SNP genotyping was performed using Affymetrix 5.0 BeadChip. Individuals with high heterozygosity or imbreeding and missingness $>0.03$ were removed and only SNPs with call rate $>0.95, \mathrm{HWE} \mathrm{P}>10^{-6}$ and $\mathrm{MAF}>0.01$ were used in the imputation.

Data Imputation and Analysis: Imputation was carried out on 413,905 SNPs using MACH 1.0 software and HapMap2 CEU (Release 22, Build 36) as reference panel. After imputation, we used SNPs with an Rsq threshold of $>0.3$ and MAF threshold of $>1 \%$. The association statistic for each SNP was calculated assuming an additive genetic model adjusting for sex, and age using a GEE model to take into account familial dependencies (GWAF R package).

## Northern Finland Birth Cohort of 1966:

Subjects: The NFBC1966 study ${ }^{9}$ [http://kelo.oulu.fi/NFBC/] includes 12,058 live born individuals, of European descent, with expected dates of birth during 1966 in the provinces of Oulu and Lapland, in Finland. Cohort has been followed up since early pregnancy until adult age. All those living in Northern Finland or in the capital area were invited to a clinical examination and blood sampling at age 31 years. Information on asthma used in this analysis was collected at this time point; 364 asthmatics and 3,502 controls were included in TAGC study.

Genotyping and QC: Genotyping was carried out using Illumina's HumanCNV370-Duo BeadChip. Individuals with high heterozygosity or imbreeding were removed and only SNPs with call rate>0.95, HWE $P>10^{-4}$ and MAF>0.01 were used in the imputation.

Data imputation and analysis: Imputation was performed on 328,007 SNPs using IMPUTE 0.3.1 and HapMap2 CEU (Release 21, Build 35). After imputation, we used SNPs with an information threshold of $>0.4$ and MAF threshold of $>1 \%$. The association statistic for each SNP was calculated assuming an additive genetic model adjusting for sex, and ancestry-informative principal components. Association analysis performed in QUICKTEST v0.94.

## EPIC-Norfolk:

Subjects: EPIC-Norfolk is part of the large multi-centre Europe-wide EPIC programme looking at the connection between diet and cancer ${ }^{10}$. There were over 30,000 participants aged $45-70$ at recruitment who lived in Norwich and the surrounding towns and rural areas. They have been contributing information
about their diet, lifestyle and health through questionnaires, and through health checks carried out by EPIC nurses. In 2006, a GWAS for obesity was carried out on 3,867 individuals. A case-cohort design was used in which the subcohort $(N=2,566)$ was a random sample of the cohort at baseline and cases were part of the remaining individuals with a value of BMI being $30 \mathrm{~kg} / \mathrm{m} 2$ or greater ( $\mathrm{N}=1,301$ ). Unlike controls in the commonly used case-control design, the subcohort was an unselected population sample allowing for a variety of traits to be investigated. The EPIC-Norfolk obese cases and the EPIC-Norfolk population based subcohorts were considered as two different samples in in the present TAGC study. Asthmatics were defined as a positive response to the question "Has the doctor ever told you that you have asthma?" which was asked at baseline survey. The remaining individuals served as healthy controls unless they had bronchitis or FEV1 < $70 \%$ predicted for age, height and sex.

Genotyping and QC: Genotyping was carried out using Affymetrix 500K BeadChip. Individuals with high heterozygosity or imbreeding and missingness $>0.06$ were removed and only SNPs with call rate>0.90, HWE $P>10^{-6}$ and MAF $>0.01$ were used in the imputation.

Data imputation and analysis: Imputation was performed on 397,438 SNPs using IMPUTE 0.3.1 and HapMap2 CEU (Release 21) as reference panel. After imputation, we used SNPs with an information threshold of $>0.4$ and MAF threshold of $>1 \%$. Data analysis was conducted separately in EPIC-Norforlk Obese and EPIC-Norforlk population-based studies (number of cases and controls are show in Supplementary Table 1). The association statistic for each SNP was calculated assuming an additive genetic model adjusting for sex, ancestry-informative principal components, and age. Association analysis between individual SNP and asthma was performed in each sub-study using SNPTEST 1.1.5.

## FINRISK

Subjects: This study is a population survey of risk factors for chronic diseases in Finland ${ }^{11}$. The survey has been executed every five years from 1972 using independent, random and representative population samples from five geographical areas of the country. Participants have completed a health-related questionnaire and undergone a physical examination including measurement of anthropometric traits and blood draw. For the FINRISK 1992 cohort, asthmatics were individuals who have answered "YES" to the question "Have you had any of the following diseases diagnosed or treated by a doctor during the past year (last 12 months)?: Asthma of the lungs". For the FINRISK 1997, 2002 and 2007 cohorts, asthmatics were individuals who have answered "YES to the question: "Have you ever been diagnosed with asthma?". In all FINRISK surveys the rest of the participants served as controls if their age was less than or equal to 70 years and they did not report pulmonary emphysema, or chronic bronchitis during the last 12 months.

Genotyping and QC: Participants were genotyped using Illumina 610K and Affymetrix 6.0 arrays. For QC, Individuals with high heterozygosity or imbreeding and missingness $>0.05$ were removed and only SNPs with call rate $>0.95, \mathrm{HWE} \mathrm{P}>10^{-6}$ and $\mathrm{MAF}>0.01$ were used in the imputation

Data imputation: Imputation was performed on 554,988 (Illumina) and 727,478 (Affymetrics) SNPs using MACH 1.0.16 and HapMap2 CEU (Release 22, Build 36) as reference panel. After imputation, we used SNPs with an Rsq threshold of $>0.3$ and MAF threshold of $>1 \%$.

## Health 2000 Study

Subjects: The study was conducted in $2000^{12}$ and included home interview, completion of several questionnaires, laboratory and anthropometrical measurements, spirometry with bronchodilator test and clinical examination by a physician. Further information was obtained by record linkage with the National Hospital Discharge Register and the National Social Insurance Institutions register data on reimbursement of asthma medication. Information about asthma was based on the following question: "Has a doctor ever diagnosed you with one of the following illnesses?" One of the listed illnesses was asthma and those who responded "YES" were considered to have asthma. The rest of the participants were taken as controls if their age was less than or equal to 70 years and they never had had chronic bronchitis.

Genotyping and QC: Samples were genotyped with Illumina 610 K or 370 K arrays. Individuals with high heterozygosity or imbreeding and missingness $>0.05$ were removed and only SNPs with call rate>0.95, HWE $P>10^{-6}$ and MAF $>0.01$ were used in the imputation

Data imputation: Imputation was performed on 555,388 SNPs using MACH 1.0.16 and HapMap2 CEU (Release 22, Build 36) as reference panel. After imputation, we used SNPs with an Rsq threshold of $>0.3$ and MAF threshold of $>1 \%$.

## Helsinki Birth Cohort Study (HBCS)

The HBCS ${ }^{13}$ includes 8,760 subjects born in Helsinki between 1934 and 1944. Between 2001 and 2004, a representative subset of 928 males and 1,075 females participated in a clinical study focusing upon cardiovascular and metabolic outcomes and cognitive functions. Information on asthma, smoking and alcohol intake is available from questionnaires for 2,003 individuals who participated in the clinical study. Information about asthma was based on the following question: "Have you ever had any of the following illnesses diagnosed or treated by a doctor?" One of the listed illnesses was "Asthma of the lungs" and those responding "YES" to this item were considered as having asthma. Those participants who responded to the same question as having emphysema or chronic bronchitis were excluded and the others were taken as controls. All participants were less than 70 years of age.

Genotyping and QC: Genotyping was performed with the Illumina 670K array. Individuals with high heterozygosity or imbreeding and missingness $>0.05$ were removed and only SNPs with call rate>0.95, HWE $P>10^{-6}$ and MAF $>0.01$ were used in the imputation

Data imputation: Imputation was performed on 546,814 SNPs using MACH 1.0.16 and HapMap2 CEU (Release 22, Build 36) as reference panel After imputation, we used SNPs with an Rsq threshold of $>0.3$ and MAF threshold of $>1 \%$.

## Young Finns Study (YFS)

Subjects: The Young Finns cohort ${ }^{14}$ is a longitudinal population study sample on the evolution of cardiovascular risk factors from childhood to adulthood. The first cross-sectional survey was conducted in 1980 in five Finnish university cities and included 3,596 participants who were in the age groups of $3,6,9$, 12,15 , and 18 years and were randomly chosen from the national population register; equal ratios of males and females were selected in each age group. In 2007, 2,204 subjects now aged 30 to 45 years participated in the latest follow-up study. Information about asthma was based on a following question: "Do you have at the moment or have you had a long-term illness, handicap or injury diagnosed by a doctor?" Those responding "YES" and specifying among the given alternatives "Asthma of the lungs" were considered as having asthma. The rest of the participants were taken as controls, except those who reported having chronic bronchitis. All YFS participants were less than 70 years of age.

Genotyping and QC: 1,963 individuals with available physician-diagnosed asthma status were genotyped with the Illumina 670K array. Individuals with high heterozygosity or imbreeding and missingness $>0.05$ were removed and only SNPs with call rate $>0.95$, HWE P $>10^{-6}$ and MAF $>0.01$ were used in the imputation Data imputation: Imputation was performed on 546,677 SNPs using MACH 1.0 and HapMap2 CEU (Release 22, Build 36) as reference panel. After imputation, we used SNPs with an Rsq threshold of $>0.3$ and MAF threshold of $>1 \%$.

Statistical analysis of FINRISK, Health 2000, HBCS and YFS: The four studies were analyzed together with an adjustment term for cohort. Association analysis between individual SNP and asthma was performed using Plink v1.07. Lambda for asthma analysis was 1.02.

## CARe Consortium

Subjects: The CARe consortium datasets which contributed to TAGC included African-American participants from four population-basoed studies: Atherosclerosis Risk in Communities (ARIC), Coronary Artery Risk Development in young Adults (CARDIA), Jackson Heart Study (JHS), and Multi- Ethnic Study of

Atherosclerosis (MESA). The number of asthma cases and controls from each study are shown in Supplementary Table 1.

CARDIA: The CARDIA study is a prospective, multi-center investigation of the natural history and etiology of cardiovascular disease in 5,115 individuals age 18 to 30 years of age at the time of initial examination (1985-1986) and drawn from four communities: Birmingham, AL; Chicago, IL; Minneapolis, MN; and Oakland, CA. Each participant's age, race, and sex were self-reported during the recruitment phase and verified during the baseline clinic visit. Genotype data were available on 955 African-American individuals. A subset of these subjects contributed to the asthma GWAS.

JHS: The Jackson Heart Study is a prospective population-based study to evaluate common complex diseases among 5,301 African Americans age 34 to 84 years at the time of initial examination (2000-2004) and drawn from the Jackson, Mississippi metropolitan area. Genotype data were available on 3,030 African-American individuals (some JHS participants are also enrolled in ARIC, and were analyzed with the ARIC dataset; 2,145 individuals are uniquely associated with JHS). Information on asthma was available for 2,162 African American individuals.

The ARIC and MESA study contains subject from African ancestry and subject of European ancestry and are described below.

In the CARe Consortium, asthma was defined as a positive response to the question "Has a doctor ever told you that you have asthma?" or "Have you ever had asthma?" on any of the available questionnaires derived from the individual studies. The agreement between these two different questions, where available to analyze, was very good, leading us to think that subject reports of asthma stemmed from of a medical diagnosis (kappa statistic with 95\% confidence intervals for individual studies are as follows: ARIC $0.82 \mathrm{Cl} 0.80-0.85$, CARDIA $0.81 \mathrm{CI} 0.77-0.83$, JHS $0.80 \mathrm{Cl} 0.78-0.83$ ). The remaining subjects served as healthy controls if they did not report any of the following conditions: wheezing, chronic obstructive pulmonary disease, emphysema, chronic bronchitis, chronic cough, chronic sputum production, or other lung disease. Individuals were excluded from the control group if they had <70 \% of the predicted FEV1 based on race- and sex-specific NHANES III prediction equations adjusting for age and height, or FEV1/FVC less than lower limits of normal for age, race and sex ${ }^{15}$.

Genotyping and QC: Samples were genotyped at the Broad Institute using the Affymetrix Genome-Wide Human SNP Array 6.0. A subset of 24 markers including a gender confirmation assay were also genotyped using the Sequenom MassArray System. Genotypes were called using Birdseed v1.33. Quality control steps were performed using the software PLINK, EIGENSTRAT and PREST-Plus (http://fisher.utstat.toronto.edu/sun/Software/Prest/). Multiple QC procedures were performed
including: confirming genotype concordance between Suquenom iPLEX and Affy6.0; removing samples with a genome-wide genotyping success rate <95\%, SNPs with genotyping success rate <90\%, monomorphic SNPs, and SNPs that mapped to several genomic locations; removing poor quality DNA (identified by estimating heterozygosity rates); removing sample duplicates, contaminated samples, and cryptic relationships (identified by using genome-wide genotype data to estimate identity-by-descent between all pairwise combinations); outlier samples were removed (identified based on nearest neighbor and "clustering based on missingness" analyses in PLINK); removing SNPs with minor allele frequency (MAF) $<1 \%$ or with genotyping success rate $<95 \%$; in JHS, excluding SNPs with an unusually high number of Mendel errors; and excluding SNPs that showed association with specific chemistry plates. Because several different ethnic groups were represented, with the expectation of differing genotype frequencies and admixture, no filters were applied for Hardy- Weinberg probability values.

Data Imputation and Analysis: To increase coverage and facilitate comparison with other datasets, we imputed genotype data using MACH version 1.0.16 ${ }^{16}$. Imputation were done using European and African ancestry reference panels. The resulting set of SNPs was conservatively filtered to remove low imputation quality ( $\mathrm{RSQ}<0.4$ ) or minor allele frequency SNPs (MAF <0.01), for which association results would be less reliable. Analysis of association of asthma with individuals SNPs was carried out using logistic regression statistical framework in PLINK for unrelated cohorts or using R (GWAF package) scripts that model family structure for family data.

## MESA: Multi-Ethnic Study of Atherosclerosis

Subjects: The Multi-Ethnic Study of Atherosclerosis (MESA) is a population-based longitudinal study of subclinical cardiovascular disease ${ }^{17}$. Between 2000 and 2002, MESA recruited 6,814 men and women 4584 years of age from six US sites that were free of clinical cardiovascular disease. The MESA Family Study recruited 1,595 African American and Hispanic family members ages 45-84 years of age specifically for genetic analysis and the MESA Air Pollution Study recruited an additional 257 participants ${ }^{18}$. Endpoints for asthma included self-reported history of asthma ("Have you ever had asthma?") and childhood asthma (Participants were asked at what age they "first developed symptoms." Those who reported symptoms at $<16$ years of age were classified as having childhood asthma). For the current effort, genetic analyses in MESA were restricted to self-reported European-ancestry participants (MESA African-ancestry were analyzed separately through CARe). Analyses of asthma included 267 cases and 2,381 controls, while analyses of pediatric asthma included 84 cases and 1361 controls.

Genotyping and QC: All participants were genotyped using the Affymetrix Human SNP array 6.0 (Affymetrix Inc., Santa Clara, CA), with 897,981 single-nucleotide polymorphisms (SNPs) passing studyspecific quality control.

Data Imputation and Analysis: IMPUTE2 was used to perform imputation of an additional $\sim 2$ million SNPs in each race/ethnic group (using HapMap Phase I and II as reference panel (release \#24, NCBI Build 36) for inclusion in GWAS analyses ${ }^{19,20}$. Multivariate additive genetic models were adjusted for age, sex, study site and principal components of ancestry. We examined genomic control values of all GWAS for evidence of residual population stratification, undetected family structure, or other sources of inflation in type I error.

## CHARGE Consortium studies included in the TAGC Consortium

## ARIC: Atherosclerosis Risk in Communities study

The Atherosclerosis Risk in Communities (ARIC) Study, sponsored by the National Heart, Lung and Blood Institute (NHLBI), is a prospective epidemiologic study conducted in four U.S. communities. ${ }^{21}$ The four communities are Forsyth County, NC; Jackson, MS; the northwest suburbs of Minneapolis, MN; and Washington County, MD. Three cohorts represent the ethnic mix of their communities; the Jackson cohort, its African-American population. ARIC was designed to investigate the etiology and natural history of atherosclerosis, the etiology of clinical atherosclerotic diseases, and variation in cardiovascular risk factors, medical care and disease by race, gender, location, and date. ARIC has coordinating, ultrasound, pulmonary, and electrocardiographic centers and three central laboratories.

The Cohort began in 1987: each ARIC field center randomly selected and recruited a cohort sample of approximately 4,000 individuals aged 45-64 from a defined population in their community. A total of 15,792 participants received an extensive examination, including medical, social, and demographic data. These participants were examined with the baseline visit occurring in 1987-89, the second visit in 199092, the third visit in 1993-95, the fourth visit in 1996-98, and the fifth visit in 2011-13. Follow-up occurs yearly by telephone to maintain contact with participants and to assess health status of the cohort. Asthma was defined based on self-report of whether the participant had been diagnosed with asthma

Genotyping and QC: Genotyping was done using the AffymetrixGeneChip SNP Array 6.0. Quality control steps for genotyping data included exclusions for call rate <95\% (participant level and SNP level), minor allele frequency $<1 \%$, HWE $\mathrm{P}<10-6$, no chromosomal location, suspected first-degree relative of an included individual based on genotype data, or more than 8 standard deviations for any of the first ten principal components. There were 9,173 Caucasian subjects with genotyping data, asthma definition, and complete covariate information.

Data Imputation and Analysis: MACH (version 1.0.16) was used to impute all autosomal SNPs with reference to HapMap CEU (release 22, build 35 ) from the 669,450 SNPs. Imputed SNPs failing additional quality control criteria (monomorphism, HWE $P<10^{-6}$, or genotype frequencies between two genotyping phases differing by $\mathrm{P}<10^{-6}$ ) were excluded, leaving $2,515,866$ genotyped or imputed SNPs for analysis. Association analysis between each individual SNP and asthma was done using logistic regression and the ProbABEL v.0.1-3 software.

## CHS: Cardiovascular Health Study

Subjects: The Cardiovascular Health Study (CHS) is a population-based cohort study of risk factors for CHD and stroke in adults $>65$ years conducted across four field centers. The original predominantly European ancestry cohort of 5,201 persons was recruited in 1989-1990 from random samples of the Medicare eligibility lists and an additional 687 African-Americans were enrolled subsequently for a total sample of 5,888 . DNA was extracted from blood samples drawn on all participants at their baseline examination. In 2007-2008, genotyping was performed at the General Clinical Research Center's Phenotyping/Genotyping Laboratory at Cedars-Sinai on 3,980 Cardiovascular Health Study participants who were free of CVD at baseline, consented to genetic testing, and had DNA available for genotyping. Participants were excluded from analysis for sex mismatch, discordance with prior genotyping, or call rate $<95 \%$. Non-European ancestry participants were excluded from this study. A total of 1,908 persons were excluded from the GWAS study sample due to the presence at study baseline of coronary heart disease, congestive heart failure, peripheral vascular disease, valvular heart disease, stroke or transient ischemic attack or lack of available DNA. Genotyping has been attempted to date in 3,397 self-identified white participants, and was successful in 3,291 persons. Of these, 3,237 subjects had information on asthma status: 179 self-reported a physician diagnosis of asthma (ever asthma), and the remaining 3,058 were used as controls.

Genotyping and QC: Genotyping was performed at the General Clinical Research Center's Phenotyping/Genotyping Laboratory at Cedars-Sinai using the Illumina 370CNV BeadChip system. Genotypes were called using the Illumina BeadStudio software as above. A set of 306,655 autosomal genotyped SNPs remained after exclusions for: call rate $<97 \%$, HWE $\mathrm{P}<10^{-5}$, more than two duplicate errors or Mendelian inconsistency (for reference HapMap CEU) or no mapping in dbSNP.

Data Imputation and Analysis: Imputation was performed using BIMBAM (v0.99) ${ }^{22}$ with reference to HapMap CEU (release 22, build 36). The analysis dataset included 2,397,023 genotyped or imputed SNPs. Statistical analysis was performed using logistic regression with robust SE estimates in $\mathrm{R}^{23}$. Adjustments were made for age at baseline, gender, and clinic site.

## deCODE Genetics

Subjects: Icelandic asthma patients over 18 years of age ( $n=1,675$ ) who attended an asthma clinic or emergency room at the National University Hospital of Iceland or the Icelandic Medical Center (Laeknasetrid) during the years 1977 to 2008 were recruited ${ }^{24,25}$. Asthma diagnosis was based on a combination of physician's diagnosis, a positive reply to the question: „Has a doctor confirmed your asthma diagnosis?", questionnaires pertaining to asthma symptoms and ICD diagnosis when receiving emergency care ${ }^{24,25}$. Atopy status (defined by at least one positive response to allergens) determined by skin prick testing was available for part of the cohort. Icelandic controls ( $n=33,408$ ) were participants from various deCODE genetics programs without known asthma. The study was funded by deCODE Genetics.

Genotyping and QC: Genome-wide SNP genotyping of 38,446 Icelanders was done using Illumina HumanHap300 or HumanHapCNV370 SNP chips. After excluding SNPs with MAF < 1\%, with genotype yield $<96 \%$, or with P value for deviation from HW equilibrium $<10^{-6}, 290,447$ SNPs were used for imputation. After excluding samples with < 98\% genotype yield, and individuals with asthma related phenotypes from the controls, 1,675 asthma cases and 33,408 controls were included in the association analysis.

Data Imputation and Analysis: Imputation of the HapMap Phase 2 Release 22 SNPs was performed using IMPUTE ${ }^{26}$ with the CEU phased haplotypes as a reference. Tests of allelic association were performed for both genotyped and imputed SNPs using logistic regression implemented in the SNPTEST software, using the -method score option to deal imputation uncertainty. To correct for relatedness of study individuals, and for possible population stratification, the resulting chi-square test statistic was divided by the genomic inflation factor $\lambda_{g}=1.26$.

## EAGLE Consortium

 COPSACSubjects: The COPSAC 2000 birth cohort study is a prospective clinical study of a birth cohort of 411 infants born to mothers with a history of asthma. The newborns were running at the age of 1 month, the recruitment of which was previously described in detail ${ }^{27-29}$. Asthma was diagnosed longitudinally by the research physicians based upon a predefined algorithm as previously described ${ }^{28,29}$. Diagnostic criteria included (1) recurrent episodes of troublesome lung symptoms recorded in the daily diary cards as five episodes within 6 months, each episode lasting at least three consecutive days; (2) symptoms typical of asthma based on doctors interviews of the parents at the clinical research unit; (3) need for intermittent rescue use of inhaled $\beta 2$-agonist; and (4) response to a 3 -month course of inhaled corticosteroids and relapse when stopping treatment. Cases in the current study were defined from having an asthma diagnosis before age 7 years and controls as children with clinical follow-up to age 7 years and no asthma diagnosis.

Genotyping and QC: High throughput genome-wide SNP genotyping were performed using the Illumina Infinium ${ }^{\text {TM }}$ II HumanHap550 v1, v3 or quad BeadChip platform (Illumina, San Diego), at the Children's Hospital of Philadelphia's Center for Applied Genomics. All family structures were validated by means of identity-by-descent comparison of all subjects in the data set (proportion identity by descent). Sex was checked by calculating heterozygosity rates of the X chromosome and comparing with reported sex. Lowquality samples were removed based on call rate (<97\%), and low-quality markers were removed based on call rate ( $<98 \%$ removed), low minor allele frequency ( $\leq 0.1 \%$ excluded), or being out of Hardy-Weinberg equilibrium ( $\mathrm{P}<10^{-6}$, excluded). Also, principal component analysis-based removal of non-European samples was performed (using HapMap as reference).

Data Imputation and Analysis: Imputation to the CEU panel of HapMap release 22 was performed using MACH 1.0.12 ${ }^{30}$ using phased haplotypes as reference. The genomic build was hg18/NCBI36. Statistical association tests were performed on $2,434,240$ variants using SNPTEST v2.2.0, using a genetic additive model and taking genotype uncertainty into account.

## Danish National Birth Cohort (DNBC)

Subjects: The Danish National Birth Cohort (DNBC) is a collection of data on 92,274 pregnant women recruited between 1996 and $2002{ }^{31}$. The participating women were interviewed twice during pregnancy and the children are followed through childhood and adolescence. Children were considered to be cases for the "ever asthma" phenotype if a) they had self-reported asthma (by mother) in the DNBC
questionnaire at age 7, or b) if they had any ICD-10 diagnosis code in the chapters J45-J46 recorded in the Danish Hospital Discharge Register before age 7.

Genotyping and QC: Genotype data were available from a genome-wide association study on preterm birth nested within the DNBC ${ }^{32}$ performed with the Illumina 660 Quad chip (Illumina, Inc., San Diego, CA). The children used in the current study were all born in gestational week 37 or later. The children as well as their parents and grandparents were born in Northwestern Europe. Subjects were filtered based on < $95 \%$ call rates, and markers were filtered based on < $98 \%$ call rates, Hardy-Weinberg equilibrium p-values $<10^{-3}$, and minor allele frequencies < 0.01. Total number of asthma cases and controls passing QC was 113 and 850 , respectively, and the total number of markers passing QC was 508,059 .

Data Imputation and Analysis: Imputation of the HapMap Phase 2 Release 22 SNPs was performed using MACH version 1.0.16 ${ }^{30}$ with the CEU phased haplotypes as a reference. Tests of allelic association were performed by logistic regression using PLINK version 1.07 for genotyped SNPs and MACH2DAT version 1.0.10 for imputed SNPs.

## GENERATION R

The Generation R Study ${ }^{33}$ is a population-based prospective cohort study from fetal life onwards in Rotterdam, the Netherlands (http://www.generationr.nl). Assessments in pregnant women and children consisted of physical examinations, fetal ultrasounds, biological samples, and questionnaires. All children were born between April 2002 and January 2006. For the current study, physician-diagnosed ever asthma (no; yes) was assessed using questions adapted from the International Study on Asthma and Allergy in Childhood (ISAAC) at age 6 years.

Genotyping and QC: A GWA scan was performed on DNA isolated from cord blood samples using the Illumina 610 Quad array. Quality control comprised exclusion of duplicates, sex mismatches and low sample call rates. Further, SNPs with call rates $<98 \%$, low minor allele frequencies ( $<0.1 \%$ ), and significant deviations from Hardy-Weinberg equilibrium ( $\mathrm{P}<1 \times 10^{-6}$ ) were excluded.

Data Imputation and analysis: Imputation was performed using MACH with HapMap2 CEU as the reference population. Statistical analyses were performed using GRIMP ${ }^{34}$.

## GINA Plus/LISA Plus

Subjects: The influence of Life-style factors on the development of the Immune System and Allergies in East and West Germany PLUS the influence of traffic emissions and genetics (LISAplus) Study is a population based birth cohort study. A total of 3,097 healthy, fullterm neonates were recruited between

1997 and 1999 in Munich, Leipzig, Wesel and Bad Honnef. The participants were not pre-selected based on family history of allergic diseases ${ }^{35,36}$.

A total of 5,991 mothers and their newborns were recruited into the German Infant study on the influence of Nutrition Intervention PLUS environmental and genetic influences on allergy development (GINIplus) between September 1995 and June 1998 in Munich and Wesel. Infants with at least one allergic parent and/or sibling were allocated to the interventional study arm investigating the effect of different hydrolysed formulas for allergy prevention in the first year of life ${ }^{37}$. All children without a family history of allergic diseases and children whose parents did not give consent for the intervention were allocated to the non-interventional arm. Detailed descriptions of the LISAplus and GINIplus studies have been published elsewhere ${ }^{35}$ and ${ }^{37}$, respectively. Information on ever having physician-diagnosed asthma and related symptoms was collected using self-administered questionnaires completed by the parents. The questionnaires were completed at $6,12,18,24$ months, 4 and 6 years of age in the LISAplus study and 1, $2,3,4$ and 6 years in the GINIplus study asking for physician diagnosed asthma at each year of age since the previous follow-up and wheeze in the past 12 months. DNA was collected at the age 6 and 10 years. Genotyping and QC: Genotyping was performed using the Affymetrix Human SNP Array 5.0 and Affymetrix Human SNP Array 6.0. Genotypes were called using BRLMM-P algorithm (5.0), respectively BIRDSEED V2 algorithm (6.0). In each of the two data sets, criteria for exclusion of individuals were: a call rate below $95 \%$, a heterozygosity outside mean $+/-4 \mathrm{sd}$, a failure of the sex check or a failure of the similarity quality control using MDS analysis based on IBS. Criteria for exclusion of variants were: a call rate below 95\%, a MAF < 0.01 and a HWE p-value <0.01 ( $n=396$ ) / < 0.00001 ( $n=755$ ).

Data Imputation and Analysis: Imputation was carried out in IMPUTE2 using the HapMap release 22, NBCI build 36 of the CEU reference population. Genome-wide association analysis of asthma phenotypes was conducted using SNPTEST V2 regressing expected allelic dosage on case-control status.

## Manchester Asthma and Allergy Study (MAAS)

Subjects: The Manchester Asthma and Allergy Study is an unselected (i.e. population-based), birth cohort study ${ }^{38-42}$. The setting is the maternity catchment area of Wythenshawe and Stepping Hill Hospitals, comprising of 50 square miles of South Manchester and Cheshire, UK, a stable mixed urban-rural population. Cases were children with asthma ever by age 6. All pregnant women were screened for eligibility at antenatal visits (8th-10th week of pregnancy). The study was explained to the parents, and informed consent for initial questionnaires and skin prick testing was obtained. Both parents completed a questionnaire about their and their partner's history of asthma and allergic diseases and smoking habits. If the pregnant woman's partner was not present at the antenatal clinic visit, an invitation was sent for
him to attend an open-access evening clinic for skin prick testing and questionnaire. Once both parents had completed questionnaires and skin prick testing, a full explanation of the proposed future follow-up for the child was given. Of the 1,499 couples who met the inclusion criteria (<10 weeks of pregnancy, maternal age >18 years, questionnaire and skin test data available for both parents), 288 declined to take part in the study. A total of 1,185 participants had at least some evaluable data. The children have been followed prospectively, and attended review clinics at ages $1,3,5,8$ and 11 years.

Genotyping and QC: DNA samples were genotyping on an Illumina 610 quad chip. The Illumina genotypes were called using the Illumina GenCall application following the manufacturer's instructions. Quality control criteria for samples included: $97 \%$ call rate, exclusion of samples with an outlier autosomal heterozygosity (scree-plot visualization) gender validation and sequenome genotype concordance. Quality control criteria for SNPs included a $95 \%$ call rate, HWE $>5.9 \times 10^{-7}$, minor allele frequency $>0.005$.

Data Imputation and Analysis: Genotypes were imputed with IMPUTE ${ }^{2}$ version 2.1 .2 with 1000 genomes and hapmap phase 3 reference genotypes. Association analysis was carried out using SNPTEST v2.4.0 ${ }^{26}$ using frequentist with the score method.

## RAINE

Subjects: The Western Australian Pregnancy Cohort (Raine) Study is a prospective pregnancy cohort where 2,900 were recruited from King Edward Memorial Hospital between 1989 and 199143. Data were collected throughout pregnancy and the children have been followed-up at ages $1,2,3,5,8,10,14,17,18,20$ and 22. Individuals who were ever diagnosed by a doctor with asthma were cases, all other individuals were classified as controls.

Genotyping and QC: Genome-wide SNP genotyping of 1,593 individuals was performed on Illumina's Human660W-Quad BeadChip. Replicates were investigated and the sample with the higher proportion of missing data was excluded. Possible cryptic relatedness between individuals in the cohort was examined, and any pair of individuals who were related with a $\pi>0.1875$ was investigated and the individual with the higher proportion of missing data was excluded. Subjects were further filtered based on < 97\% call rates. SNP QC was carried out in accordance to WTCCC recommendations ${ }^{44}$; in short, markers were filtered based on $<95 \%$ call rates, Hardy-Weinberg equilibrium p-values $<5.7 \times 10^{-7}$, and minor allele frequencies < 0.01 . The total number of markers passing QC was 535,632 .

Data Imputation and Analysis: Imputation was performed using MACH version 1.0.18c and Minimac with the HapMap2 release 22 as a reference. Tests of allelic association were performed for both genotyped and imputed SNPs using logistic regression with SNPTEST v2.2.0.

## B58C: British 1958 birth cohort

Subjects: The British 1958 Birth Cohort (also known as the National Child Development Study) is based on all persons born in Britain during one week in 1958. They were followed up at ages 7,11 and 16 years by parental interviews and examinations by school medical officers, and at ages 23,33 and 42 years by means of interviews. The presence of diagnosed asthma or a history of wheezy bronchitis were ascertained by parental interviews in childhood ${ }^{45}$. In adulthood, questionnaires were used to define asthma ever, wheezing ever, and wheezing episodes in the previous 12 months ${ }^{46,47}$. During 2002-2004 all cohort members still in contact with the study team were visited for a biomedical examination ${ }^{48}$ and invited to contribute blood samples with consent for DNA extraction. The lifetime prevalence of asthma or wheeze in the full cohort was $49 \cdot 9 \%$ and in the subgroup with DNA samples was $48.8 \%$.

For the purposes of this meta-analysis, cases ( $\mathrm{N}=986$ ) were defined as those with a history of "asthma ever" at any age up to 42 years, and controls ( $\mathrm{N}=5,505$ ) were defined as the remainder of the cohort, including those with a history of wheeze or wheezy bronchitis, but no asthma. This followed the convention set by the GABRIEL consortium ${ }^{6}$.

Genotyping and QC: Non-overlapping subsets of the British 1958 cohort DNA collection were genotyped as control samples by the Wellcome Trust Case-Control Consortium ${ }^{49}$ and the Type 1 Diabetes Genetics Consortium ${ }^{50}$, using the Illumina 550K array. From the remainder of the samples, all asthmatic cases and a sample of non-asthmatics were genotyped by the GABRIEL consortium using the Illumina 610K array ${ }^{6}$. These three genome-wide datasets were combined before imputation, selecting 532,203 SNPs which were common to the three deposits.

Each of the three subsets was restricted to subjects of white European ethnicity, but ancestry outliers on principal components analysis were excluded from all analyses. Samples with low average call rate ( $<98 \%$ ) and sex mismatches were also removed. SNPs used for imputation excluded those with a call rate $<95 \%$, minor allele frequency $<1 \%$, HWE p-value $<0.0001$ or discordant allele frequencies across the three deposits ( $\mathrm{p}<0.0001$ for pairwise comparisons). For chromosome X, the test of HWE was restricted to females, and additionally, SNPs with discordant allele frequencies between males and females ( $p<0.0001$ ) were excluded. As a result, 494,161 (93\%) of the 532,203 genotyped SNPs were used for imputation.

Data Imputation and Analysis: Imputation against the HapMap 2 (release 21) CEU reference panel was carried out using MACH 1.0.16. Statistical analysis used the palogist procedure in ProbABEL 0.1-3 ${ }^{51}$.

## EVE Consortium

STAMPEED: SNP Typing for Association with Multiple Phenotypes from Existing Epidemiologic Data

Subjects: The STAMPEED asthma project includes subjects from the Chicago Asthma Genetics Study (C.A.G.) as well as from SARP and CSGA. CAG included both European American and African American subjects collected at The University of Chicago from a) families ascertained through affected sib pairs, b) affected children and their parents, c) adults and children with severe persistent asthma, and d) nonasthmatic control subjects (over the age of 18 years). Samples a-c were recruited in the adult and/or pediatric asthma clinics at University of Chicago Hospital; controls were recruited from the medical center at large. SARP and CSGA included both European American and African American subjects. Control subjects and subjects with mild to severe asthma were recruited at the NHLBI funded Severe Asthma Research Program (SARP) centers and the NHLBI Collaborative Studies on the Genetics of Asthma (CSGA) and characterized according to asthma severity (see ${ }^{52,53}$ ).

Genotyping and QC: Genotyping was performed on the Illumina 1Mv1 platform, with individual genotypes called using clustering algorithms as implemented in the BeadStudio software by Illumina. The resulting number of markers was $1,033,467$ prior to additional QC measures. Markers were filtered based on $95 \%$ call rates, Hardy-Weinberg equilibrium p-values $>10^{-5}$, consistency in allele frequency from the HapMap ASW (chi-square p-value $>10^{-5}$ ), and $<5$ heterozygous genotype calls in males for X -linked markers. The total number of markers following QC was 1,025,129. Subjects were filtered based on $95 \%$ call rates, matching genetic and reported sex (Fstat on X chromosome between -0.2 and 0.3 for females, and between 0.8 and 1 for males), consistency in self-reported ethnicity based on a principal component analysis in Eigenstrat (no obvious clustering with the HapMap CEU for African Americans, and the HapMap YRI for European Americans), and high or low heterozygosity (Fstat < 0.5 and > -0.2). Samples were flagged for unexpected pairwise relatedness (IBD > 30\%) or genetic identity (IBS > 90\%), with subsequent filtering performed by selecting a single sample from the pair (or group) having the highest call rate.

Data Imputation and Analysis: Imputation of the phase 2, release 21 HapMap SNPs was performed using MACH, with the consensus CEU haplotypes used as a reference for the European American cases and controls, and the combined consensus CEU and YRI haplotypes for the African American cases and controls. Tests of allelic association were performed on dosages for both genotyped and imputed SNPs using logistic regression in R (http://CRAN.R-project.org/). For the set of African American cases and controls, local ancestry was used as a covariate at the SNP level as estimated using the set of genotyped SNPs in the program LAMP - Local Ancestry in adMixed Populations ${ }^{54}$. Admixture in African Americans was modeled under 7 generations of admixture with a 2-population model of $81 \%$ ancestry from Africa and $19 \%$ ancestry from Europe ( $\alpha=[0.81,0.19]$ ) as estimated based on the first principal component in a PCA analysis using EIGENSTRAT ${ }^{55}$ ). Windows were offset by a factor of 0.2 , the cutoff for linkage was set to 0.1 ,
and a constant recombination rate was set to $10^{-8}$. Local ancestry for imputed SNPs was obtained from the next closest genotyped SNP.

## CHS: The Children's Health Study

Subjects: The Children's Health Study (CHS) is an ongoing cohort study in southern California investigating both genetic and environmental factors related to childhood asthma and lung function growth. The Children's Health Study GWAS was based on a nested case-control sample of 1,249 asthmatics and 1,751 controls selected from within the cohort. All subjects in this GWAS sample were either Hispanic white (HW) ( $n=1,398$ ) or non-Hispanic white (NHW) ( $n=1,602$ ). Based on questionnaire responses, children were characterized as having doctor-diagnosed asthma at study entry or during active follow-up (cases), or as never having a diagnosis of asthma (controls).

Genotyping and QC: Study samples were genotyped at the USC Epigenome Center using the Illumina HumanHap550, HumanHap550-Duo or Human610-Quad BeadChip microarrays. Individuals were excluded from analysis with call rates < 90\% ( $\mathrm{n}=155$ ). The HumanHap550, HumanHap550-Duo and Human610-Quad contained 366,366 and 418 SNPs respectively that overlapped with a candidate gene study containing a large number of the subjects in this study ( $\mathrm{n}=2,905$ ). The average concordance rate between matching subjects for these overlapping SNPs was $>99.69 \%$ for $>99 \%$ of the samples having a call rate $>90 \%$. Subjects with poor concordance with genotypes from the candidate gene study were excluded ( $\mathrm{n}=19$ ). SNPs were excluded prior to imputation if they were not concordant between Illumina genotyped HapMap samples and HapMap2_r21 (< 95\% in any population) or not concordant between Illumina genotyped HapMap samples on the HumanHap550 and Human610 (<95\% in any population) ( $n=8,616$ ). Additionally, SNPs were filtered if they had a call rate $<95 \%(n=84,381)$ or departed from Hardy-Weinberg equilibrium ( $\mathrm{p}<10^{-5}$ ) in controls in HW and NHW samples separately ( $\mathrm{n}=762$ and 766 respectively).

Data Imputation and Analysis: Imputation was performed with MACH v1.0.16 using the HapMap phase 2 release 21 consensus CEU haplotypes as a reference for the NHW subjects and CEU+ASN haplotypes for the HW subjects. Tests of allelic association were performed on dosages for both genotyped and imputed SNPs using logistic regression using R (www.R-project.org), adjusting for age, community of residence, sex, and ancestry covariates derived from the program STRUCTURE ${ }^{56}$ applied to 557 ancestrally informative markers.

## GRAAD: Genomic Research on Asthma in the African Diaspora

Subjects: The Genomic Research on Asthma in the African Diaspora study consists of 498 asthma cases and 500 non-asthmatic controls from the Baltimore-Washington, D.C. metropolitan area who self-
reported as African American. Because asthma is often characterized by onset during childhood, there was a deliberate decision to favor adults in the control group to minimize including controls with the potential for developing asthma. A standardized questionnaire based on either the American Thoracic Society ${ }^{57}$ or International Study of Asthma and Allergy in Childhood (ISAAC) ${ }^{58}$ was administered by a clinical coordinator. Asthma was defined as both a reported history of asthma, and a documented history of physician-diagnosed asthma (past or current). All controls (except 50, see below) were likewise administered a standardized questionnaire and were determined to be negative for a history of asthma. Asthma status on 50 controls participating in a study of the genetics of human pigmentation ${ }^{59}$ was not explicitly determined, although "known clinical disease" was among the exclusion criteria. The study protocol was approved by the institutional review board at either the Johns Hopkins University or Howard University.

Genotyping and QC: Details of genotyping and quality control (QC) have been previously described ${ }^{60}$. Briefly, genotypes were generated at the Center for Inherited Disease Research (CIDR) for 665,352 SNPs on the Illumina HumanHap650Y Versions 1 and 3 BeadChips and the Illumina Infinium II assay protocol. Genotypes were released for 994 GRAAD samples for SNPs with < 5\% missing data, no replicate error among other standard QC protocols. Relationships between individuals within each study were verified using PLINK and RELPAL ${ }^{61}$, and marker-level QC parameters (including minor allele frequency, differential missing rates between cases and controls, and Hardy Weinberg Equilibrium) were evaluated in PLINK. The genetic structure of African American cases and controls was evaluated using unrelated individuals from the three HapMap "continental" ancestral populations (CEU, YRI, and ASN; www.hapmap.org) using 416 SNPs identified as ancestry informative markers (AIMs) selected for maximal difference between African and European populations in STRUCTURE.

Data Imputation and Analysis: Genotypes were imputed at all HapMap phase 2, release 21 SNPs using MACH, and using a combined panel of the HapMap CEU, YRI and ASN phased haplotypes. For the casecontrol sample (GRAAD), a two sample t-test was used to compare the allele dosage between cases and controls.

## SAPPHIRE: The Study of Asthma Phenotypes and Pharmacogenomic Interactions by Race-Ethnicity

Subjects: Study participants received their care through a large health system serving southeast Michigan. Cases were part of the Study of Asthma Phenotypes and Pharmacogenomic Interactions by Race-Ethnicity (SAPPHIRE) ( $\mathrm{n}=149$ ), and met the following criteria: age 12-56 years, a prior clinical diagnosis of asthma, no recorded diagnosis of chronic obstructive pulmonary disease (COPD) or congestive heart failure (CHF), a baseline $\mathrm{FEV}_{1}$ between $40-90 \%$ predicted, $>12 \%$ baseline bronchodilator reversibility, no smoking in the
preceding year or <10 pack-year smoking history total, no oral or inhaled corticosteroid use in the 4 weeks preceding screening, and not pregnant at the time of enrollment and not intending to get pregnant during the study period. Controls were obtained from a separate cohort, the Wayne County Health Environment Allergy and Asthma Longitudinal Study. All were women $\geq 21$ years of age who also resided in the Detroit metropolitan area and received their care from the same health system. The analytic group for the metaanalysis was restricted to individuals who reported being African American. The Institutional Review Board of Henry Ford Health System approved all of the components of this study.

Genotyping and QC: Genotyping for the cases was performed on the Affymetrix Genome-Wide Human SNP Array 6.0, whereas control individuals were genotyped on either the Mapping 500K Array and the Genome-Wide Human SNP Array 5.0 (Affymetrix, Inc., Santa Clara, California). Genotyping calls and the chip quality control (QC) call rate were assessed using Affymetrix Genotyping Console. Subjects with the following were excluded from the analysis: missing information for one chip (i.e., for those genotyped on the 2 chip Mapping 500K Array), genetic sex inconsistent with reported sex, chips which did not meet the manufacture's recommended QC call rate, and $<90 \%$ overall call rate. In order to appropriately match cases and controls, genetic ancestry was estimated in all individuals using markers informative for African and European ancestry and the program PSMIX ${ }^{62}$. We restricted the control set to individuals with $\geq 30 \%$ African ancestry ( $\mathrm{n}=132$ ), which was the lower limit of African ancestry among the cases. Single nucleotide polymorphisms (SNPs) common to all genotyping platforms were selected and subjected to the following additional criteria: call rate $\geq 95 \%$ and an exact Hardy-Weinberg equilibrium test with $p$-value $>10^{-5}$ among the controls.

Data Imputation and Analysis: A total of 404,088 SNPs were used to impute the HapMap phase 2, release 21 SNPs using MACH with the phased HapMap CEU and YRI haplotypes as a reference. Case/control association tests for asthma status were performed using logistic regression in $R$ (http://CRAN.Rproject.org/) on genotype dosages, and adjusting for the first principal component from EIGENSTRAT.

## JAARC: Japanese Adult Asthma Research Consortium

Subjects: We selected subjects with childhood-onset asthma from the case subjects who participated in a genome-wide association study (GWAS) for adult asthma in the Japanese population ${ }^{63}$. A total of 301 subjects with a history of childhood-onset asthma (asthma onset by 15 years of age; age [mean $\pm \mathrm{sd}$ ], $39.4 \pm 14.8$ ) years; female [\%], 47.8) were recruited by physicians' interviews. All subjects with asthma were diagnosed by physicians according to the American Thoracic Society criteria. A total of 3,304 control subjects for the GWAS (age [mean $\pm$ sd], $52.5 \pm 15.1$ ) years; female [\%], 44.4) were recruited as described ${ }^{63}$.

Genotyping and QC: For the GWAS, we genotyped 301 cases using the Illumina HumanHap610-Quad BeadChip and 3,304 controls using the Illumina HumanHap550 BeadChip. We conducted allele sharing analysis and performed principal component analysis for the genotype data of the samples along with European (CEU), African (YRI) and East Asian (Japanese [JPT], and Han Chinese [CHB]) individuals obtained from the Phase II HapMap database by using SMARTPCA. We excluded related samples and outliers who were away from the east Asian cluster. We also excluded SNPs with minor allele frequencies of less than 0.01 in both cases and controls. SNPs having call rates $\geq 99 \%$ in both cases and controls were used for the association study. We conducted exact Hardy-Weinberg equilibrium (HWE) analysis, and SNPs with $P$ values less than the cut-off values of the HWE test ( $P<10^{-6}$ in controls) were also excluded from the analysis.

Data Imputation and Analysis: Imputation of HapMap2, Release 21 SNPs was performed using MACH with the JPT+CHB haplotypes as a reference. SNPs with $P$ values of HWE in the control group $\geq 1 \times 10^{-6}$ were imputed. In the GWAS, the statistical significance of the association with each SNP was assessed using a logistic regression analysis.

## JPAC: Japan Pediatric Asthma Consortium

Subjects: The part of the study population, genotyping and quality control have been previously described ${ }^{64}$. All subjects with asthma were child or child-onset (<15 years old) asthmatics in Japan. Patients were recruited from 3 pediatric hospitals and 1 pediatric clinic, and the diagnosis of the asthma in all patients was confirmed by specialists in pediatric allergology on the basis of the criteria of the National Institutes of Health, USA, with minor modifications. The control for the GWAS were healthy Japanese adult subjects from Tokyo who had no current history of asthma. All subjects were self-reported Japanese origin. Genotyping and QC: Genome-wide SNP genotyping of 978 pediatric asthmatics and 694 controls was performed on Illumina's HumanHap550v3/610 Genotyping BeadChip (Illumina, Inc., San Diego, CA). We examined the potential genetic relatedness on the basis of pairwise identity by state for all of the successfully genotyped samples using the EIGENSTRAT software, and we excluded samples belonging to Han Chinese or Ryukyu, an island located in southern Japan. Subjects were filtered based on < 95\% call rates, and markers were filtered based on < 95\% call rates, Hardy-Weinberg equilibrium p-values < $10^{-4}$, and minor allele frequencies $<0.01$. Total number of pediatric asthmatics and controls passing QC was 938 and 672, respectively, and the total number of markers passing QC was 482,803 for HumanHap 610 and 435,199 for HumanHap550v3.

Data Imputation and Analysis: Imputation of the HapMap Phase 2 Release 21 SNPs was performed using MACH version 1.0.17 with the JPT+CHB phased haplotypes as a reference. Tests of allelic association were performed for both genotyped and imputed SNPs using logistic regression with MACH2DAT software version 1.1.9.

## GABRIEL Consortium

The GABRIEL Consortium includes 19 studies that contributed to TAGC. Seven of the 19 studies were split into two datasets according to age of onset of asthma (before or after 16 years of age); the INDUSTRIAL subgroup included four studies and the German MAS and MAGICS studies were combined.

EGEA: Briefly, the EGEA study ( $\mathrm{N}=2,120$ subjects) combines a case-control and a family-based study of asthma cases with three surveys over 20 years (EGEA1: 1991-1995, EGEA2: 2003-2007 and EGEA3: 20112013). The whole study population included 388 asthmatic probands recruited in chest clinics and their 1,317 family members (probands' parents and/or siblings) plus 415 population-based controls ${ }^{65}$. The probands (asthmatics and controls) were between 7 and 70 years old at time of study. All subjects were of European ancestry and were born in France. Data collected through face-to-face interviews and examination included extensive phenotypic characterization (detailed clinical data based on standardized questionnaire, skin prick tests, lung function tests, bronchial responsiveness, blood samples, white blood cell counts, total IgE), data on risk factors (environmental exposures, diet, physical activity, hormonerelated events) and drug consumption. For the present study, cases with asthma were defined by a positive answer to the question "Have you ever had asthma attacks?" at EGEA1 or EGEA2. Detailed description of the study can be found in Kauffmann et $\left.a\right|^{65,664}$ and at http://egeanet.vif.inserm.fr. After QC, a total of 1,928 EGEA subjects were retained in the GWAS analyses.

ALSPAC: is a population-based, prospective, longitudinal, birth-cohort study that was recruited during pregnancy. Pregnant women resident in Avon, United Kingdom and with estimated dates of delivery 1/4/1991-31/12/1992 were recruited through antenatal clinics. ${ }^{67}$ (www.bristol.ac.uk/alspac) Of 14,451 pregnancies recruited, there were 14,072 live births and 13,988 children remained alive at age 1 year. Children have been followed from birth using a combination of self-completion questionnaires sent at regular intervals to their mothers and hands-on assessments at annual dedicated research clinics from age 7 years. Asthma in children was defined as a positive response to the question, "Has a doctor ever told you that your child has asthma?" in a questionnaire sent to their mothers at 91 months after birth. A total of 5,231 children had data on asthma at 91 months with DNA available for genotyping. Of these, 650 (12.4\%) had asthma. An equal number of controls were selected at random from the remaining population with
questionnaire responses at 91 months and DNA available. After QC a total of 1,216 subjects were retained in the GWAS analyses.

ECRHS: Sixteen centres (eight countries) in the ECRHS contributed samples to the GWAS. ${ }^{68,69}$ (www.ecrhs.org) In each centre, a representative community-based sample of at least 3000 adults aged 20-44 years were invited to complete a brief postal questionnaire asking about respiratory symptoms (ECRHS I - Stage 1) between 1991-1993. A random sample of these ( 600 per centre) underwent intensive further investigation (ECRHS I - Stage 2 - random sample). Participants who had symptoms highly suggestive of asthma but who had not been selected at random to take part in Stage 2 were also invited to undergo intensive investigations (ECRHS I-Stage 2- enriched sample). About ten years later all adults who had taken part in Stage 2 were re-contacted (ECRHS II) and again asked about respiratory symptoms. Samples suitable for DNA extraction were collected. For the GWAS initiative all cases of asthma were identified (participants from the random or enriched sample who said yes to the question 'Have you ever had asthma?' at either ECRHS Stage I or Stage II). Controls were a random sample (of the random sample) who answered 'no' to the same question in both surveys. After quality control (QC) criteria a total of 2,210 ECRHS subjects were retained in the GWAS analyses.

BAMSE: Between 1994 and 1996, 4,089 newborn infants were recruited in the BAMSE study, and questionnaire data on baseline study characteristics were obtained. ${ }^{70,71}$ The catchment area included central and north-western parts of Stockholm. At approximately one, two, four, and eight years of age, parents completed questionnaires on their children's symptoms related to asthma and other allergic diseases. The response rates were $96 \%, 94 \%, 92 \%$ and $84 \%$, respectively. At eight years of age, all children of the BAMSE study were invited to clinical testing, and blood samples were obtained from 2,480 children. DNA was extracted from 2,033 samples after exclusion of samples with too little blood, lack of questionnaire data, or if parental consent to genetic analysis of the sample was not obtained. From these samples, all children with a doctor's diagnosis of asthma (ever) were selected as cases and children with no history of asthma or other allergic diseases were selected as controls. After QC a total of 485 subjects were retained in the GWAS analyses.

BUSSELTON: Residents of the town of Busselton in the southwest of Western Australia have been involved in a series of health surveys since 1966. (www.busseltonhealthstudy.com) The population is predominantly of European origin. In 1994/95 there was a follow-up study involving a subset of those who had attended any of the previous surveys. Cases of asthma were defined as those who reported doctordiagnosed asthma at any survey that they attend from 1966 to 1994 (answer 'Yes' to 'Has your doctor ever told you that you had asthma?'). ${ }^{72}$ Controls are those who have consistently answered 'No' to 'Has your
doctor ever told you that you had asthma?' at all previous surveys that they have attended from 1996 to 1994. For the GWA study, a case control sample of unrelated individuals was selected. After QC a total of 1,207 subjects were retained in the GWAS analyses.

GABRIEL ADVANCED SURVEYS: Are cross-sectional population-based surveys conducted in rural areas of Austria, Germany, and Switzerland. In total, 135,359 children aged 6-12 years were addressed through schools. In a first stage in fall/winter 2006, asthma, allergic disease, and contact to farming environments were assessed using a short parental questionnaire ( $n=79,888$ ). In a second stage in spring/summer 2007, 9,668 children were selected among families consenting in writing to blood sampling, genetic testing and collection of environmental samples by stratified random sampling to ensure representation of children with high exposure to farming environments. ${ }^{73}$ Genomic DNA was purified from blood samples using the Puregene chemistry (QIAGEN, Hilden, Germany) on an Autopure LS instrument (QIAGEN, Hilden, Germany). Genomic DNA and questionnaire data were available for 7,303 children of whom 862 cases and 865 controls were selected for genotyping. A case was defined as a parental report of asthma diagnosed by a doctor at least once or asthmatic bronchitis diagnosed at least twice during lifetime. To account for the stratified random sampling, probability weights were introduced in the statistical analyses. After QC a total of 1,692 subjects were retained in the GWAS analyses.

KSMU: is a population-based case-control study of adult cases of asthma and controls matched for age and sex. ${ }^{74,75} \mathrm{~A}$ total of 429 unrelated subjects were recruited in this study, ( 215 patients with asthma and 214 controls). The study subjects were of Russian origin from Central Russia. All patients were recruited at the Department of Pulmonology, Kursk Regional Clinical Hospital between 2003 and 2004. Additional adult patients with asthma and healthy subjects (>200 samples) from the same population were recruited between 2007 and 2008 specially in order to increase final sample size for the GWAS initiative. All patients were diagnosed with asthma by the presence of characteristic symptoms, reversibility of airway obstruction or airway hyperresponsiveness to methacholine. All control subjects were enrolled in accordance with the following criteria: (1) no symptoms and history of allergic diseases, (2) normal total serum IgE levels, (3) and normal pulmonary function test results. Personal data, including smoking status and age of the disease onset, was collected through in-person interviews. After QC a total of 568 subjects were retained in the GWAS analyses.

MRCA-UK: is a collection of nuclear families (207 families total number of subjects 783) ascertained through a proband with $\geq$ Step 3 asthma according to British Thoracic Society guidelines ${ }^{76}$ and four hundred and thirty seven non-asthmatic UK controls (UK-C) ${ }^{77}$. Probands and family members all answered a nurseadministered comprehensive respiratory questionnaire with asthma being defined as positive response to
the question "Has your doctor ever told you that you have asthma?". The children from this collection together with those from MAGICs (below) formed the case control dataset genotyped in the first GWAS for asthma ${ }^{77}$.

MAGICS consists of 655 asthmatic children (asthma diagnosed by a pulmonologist) were derived from the Multicenter Asthma Genetics In Childhood Study with a further 73 asthmatics and 694 unaffected individuals (randomly selected) from Phase One of the International Study of Asthma and Allergy in Childhood (ISAAC) ${ }^{78,79}$. For the 73 asthmatics from ISAAC, asthma was physician diagnosed and/or reports by parent of recurrent spastic or asthmatic bronchitis. MAGICS together with the children from the MRCA collection formed the case control dataset genotyped in the first GWAS for asthma ${ }^{77}$.

MAS: Consists of 1,314 healthy mature children born in 5 German cities in the year 1990. All children were followed at the age of $1,3,6,12,18$, and 24 months and at yearly intervals thereafter until age 15 years. ${ }^{80,81}$ Clinical and environmental assessment included standardized interviews, questionnaires, physical examinations, and environmental studies. Total and specific $\lg E$ antibodies to hen's egg, cow's milk, wheat, and soy were determined at the age of $1,2,3,5,7$, and 10 years. DNAs from all children with a doctor diagnosis of asthma were provided for the GWA study. This dataset was combined with the MAGICS dataset.

PIAMA: Is a birth cohort study consisting of two parts: a placebo controlled intervention study in which the effect of mite impermeable mattress covers was studied and a natural history study in which no intervention took place. In this study, only data from the natural history part of the study are presented. Details of the study design have been published previously. ${ }^{82-84}$ Recruitment took place in 1996-1997. A screening questionnaire was distributed to 10,232 pregnant women visiting one of 52 prenatal clinics. 10,232 pregnant women completed a validated screening questionnaire at their prenatal health care clinic. Based on this screening, 7,862 women were invited to participate, of whom 4,146 women agreed and gave informed consent. Mothers reporting a history of asthma, current hay fever or allergy to pets or house dust mite were defined as allergic. Follow-up of the children took place at 3 months of age and yearly from 1 to 8 years of age. The response rates to the annual questionnaires ranged from $3030(92 \%)$ at age 1 to 2732 ( $83 \%$ ) at age 8 years. For the GWA study, DNA from childhood asthma cases and a set of matched non-asthmatic controls were provided. After QC a total of 359 subjects were retained in the GWAS analyses.

SAPALDIA: Contributed all self-reported asthma cases as well as a random sample of controls. These subjects were obtained from among 6,055 SAPALDIA cohort subjects that participated in both, the baseline (1991) and follow-up (2002) examinations and agreed to providing blood for genetic analysis. SAPALDIA is
a population-based cohort that originally recruited subjects aged 18 to 60 from population registries in eight Swiss communities representing the three largest language groups (German, French, Italian) as well as different levels of air pollution, altitude and degrees of urbanization. ${ }^{85-87}$ At both baseline and followup examination subjects underwent spirometry as well as a detailed interview on respiratory health, smoking history, lifestyle factors and anthropometry. At follow-up, 8,047 of 9,651 baseline subjects reparticipated in at least one part of the study and a formal biobank was established. SAPALDIA questions about smoking and asthma status were equivalent to those used by the ECRHS. Asthma status was defined by an affirmative answer to the question "Have you ever had asthma" at baseline and/or follow-up interview. After QC a total of 1,521 subjects were retained in the GWAS analyses.

TOMSK: Is a population-based family study conducted by the Research Institute of Medical Genetics and Siberian State Medical University (Tomsk, Russia) from 1998 onwards. ${ }^{88,89}$ Both nuclear families and extended pedigrees were recruited through atopic bronchial asthmatic probands. All participants were Russians or of a mixed ethnic origin due to marriages between Russians and major East Slavonic populations (Ukrainians, Byelorussians). Altogether, 196 families were studied, out of which 150 families were recruited in Tomsk Region Children Hospital and Tomsk Region Hospital (Tomsk, Russia), and 46 families were recruited in the city of Irkutsk hospitals by the staff of the Irkutsk State Institute of Doctor's Advanced Training (Irkutsk, Russia). Both probands and their relatives were clinically examined to establish diagnosis of asthma and atopy by the GINA criteria (Global Initiative for Asthma: Global Strategy for Asthma Management and Prevention. http://www.ginasthma.org). Besides the clinical examination, laboratory and functional testing were conducted to assess common IgE levels (solid-phase immuneenzyme assay), specific sensitization (skin-prick tests), lung volumes (spirometry), and airway responsiveness (bronchoprovocative tests with methacholine). After QC a total of 681 subjects were retained in the GWAS analyses.

UFA: Is a population-based case-control study of asthma cases and controls matched on age and sex. Cases are unrelated patients with physician-diagnosed asthma and controls disease-free. Subjects are of different ethnic origins (Russians, Tatars and Bashkirs) from Volga-Ural region of Russian Federation. The Volga-Ural region is located at the border between Europe and Asia and has been the arena of permanent genetic exchanges among Siberian, European, Central Asian and other populations. Anthropologically, Russians, Tatars and Bashkirs are Caucasians and have a varying Asian component. Recruitment of cases and controls was carried out by the Hospital of the Bashkir Medical State University and Ufa Municipal Hospital №21 between 1999 and the year 2007. Asthma patients were diagnosed by pulmonologists on the basis of clinical examination, family and medication history, objective tests of lung function. The
controls were healthy subjects who met all the following criteria: (1) no symptoms or history of asthma or other pulmonary diseases; (2) no symptoms or history of atopy; and (3) absence of first-degree relatives with a history of asthma or atopy. After QC a total of 678 subjects were retained in the GWAS analyses. INDUSTRIAL (the Industrial Cohorts Research Group): The study base consists of several pooled surveys on occupational asthma from Denmark and The Netherlands. The populations come from industries with exposure to known major allergens and irritants; high molecular weight sensitizers (wheat flour, fungal alpha amylase, animal allergens), low molecular weight sensitizers (isocyanate monomers and oligomers) and irritants and inflammatory agents (isocyanates, organic dust, endotoxins, wood). Prevalent cases and industry, gender and age matched study-specific controls were included for the GWAS initiative. Briefly, cases and controls originate from the following population based or prospective cohort studies in agricultural dust and/or animal exposed environments: the Danish SUS study ${ }^{90,91}$, the Dutch Omega ${ }^{92,93}$ and Veterinarian's Health Study ${ }^{94}$; in wheat and amylase exposed environments: the bakers from the Dutch Bakers health surveillance project ${ }^{955}$; in wood dust exposed environments: workers in the Danish furniture industry of the Danish Wood Dust Cohort; ${ }^{96,97}$ and for isocyanate exposed workers: a population of Dutch industrial spray painters. ${ }^{98,99}$ Asthma, respiratory symptoms and potential confounders like smoking were primarily assessed from questionnaires. More objective health measures were also collated including lung function measurements, a specific bronchial hyperresponsiveness and specific $\operatorname{lgE}$ measurements. Exposure was assessed by taking job histories from questionnaires. For all studies extensive measurement series have been collected which have been used to create generic or study specific Job Exposure Matrices by which the exposure can be estimated quantitatively on the basis of job title and sometimes tasks performed ${ }^{100}$. After QC a total of 1,227 subjects were retained in the GWAS analyses.

SEVERE: Severe/refractory cases of asthma were recruited through three specialist severe asthma clinics; adult and childhood clinics based at the Royal Brompton Hospital, London and an adult clinic at the Glenfield Hospital, Leicester. Patients attending the Glenfield Hospital clinics had full characterisation and were deemed to have severe/refractory asthma according to a specialised protocol involving parameters of airway inflammation, airway physiology, as well as quality of life and control of symptoms. ${ }^{101,102}$ Those attending Royal Brompton Hospital adult clinics were also fully characterised, with severe asthma defined according to the ATS and ERS definition of severe asthma. ${ }^{103,104}$

Severe asthma in the pediatric clinic was defined as one or more of the following criteria: (1) Persistent (most days, for at least 3 months) chronic symptoms (the necessity because of symptoms for short-acting $\beta-2$ agonists at least three times/week) of airways obstruction despite high dose inhaled corticosteroids
(Beclomethasone equivalent $800 \mathrm{mcg} /$ day) and trials of every add-on medication available in the country of residence (these would include, if available, long acting $\beta-2$ agonist, leukotriene receptor antagonist, oral theophylline in the low, anti-inflammatory dose). This group includes Type 1 brittle asthma. (2) Recurrent severe asthma exacerbations despite attempts with medication including trials of allergen avoidance, low dose daily inhaled corticosteroids or intermittent high dose inhaled corticosteroids: either at least one admission to an intensive care unit, or at least two hospital admissions requiring intravenous medication/s, or $\geq 2$ courses of oral steroids during the last year, despite the above therapy. This group includes Type 2 brittle asthma. (3) Persistent airflow obstruction: post oral steroid, post-bronchodilator Z score <-1.96 for $\mathrm{FEV}_{1}$, with appropriate normative data despite the above therapy. (4) The necessity of prescription of alternate day or daily oral steroids to achieve control of asthma. Children were evaluated in detail to exclude as far as possible non-adherence to therapy, significant co-morbidity (for example, rhinosinusitis and gastroesophageal reflux), psychosocial issues and adverse environmental circumstances as contributing factors to the severity of asthma. ${ }^{104-107}$ Because of the relatively small numbers of severe asthmatics, this group were not subdivided into childhood onset and adult onset groups. After QC a total of 290 subjects were retained in the GWAS analyses.

For the SEVERE cohorts, control data for analysis was obtained through the WTCCC (http://www.wtccc.org.uk). Data used was from 974 samples genotyped using an Illumina 1.2M custom chip.

Genotyping and QC for all GABRIEL datasets: All Gabriel consortium datasets, except for the MRCA and MAGICS datasets, were genotyped at Centre National de Génotypage (CNG, Evry, France) using the Illumina Human610-Quad array ${ }^{6}$. The MRCA and MAGICS datasets were genotyped using Illumina Sentrix Human-1 and Sentrix HumanHap300 BeadChips, as part of the first asthma GWAS ${ }^{77}$. QC of individuals and SNPs genotyped at CNG was done in each dataset following the same protocol ${ }^{6}$. Briefly, individuals were removed from analysis if they were not of European descent (principal component analysis of each GABRIEL datasets with all HapMap populations), had a low genotyping call rate (<95\%) or were discrepant or ambiguous for genetic sex. Single Nucleotide Polymorphisms (SNPs) with call rate lower than 95\% or minor allele frequency (MAF) lower than 0.01 , or with Hardy-Weinberg equilibrium P-value $<10^{-4}$ were removed. QC for MRCA and MAGICS is detailed Moffatt et al. ${ }^{77}$

Imputation and analysis of all GABRIEL datasets: In each dataset, genome-wide imputations were performed using MACH 1.0 software and HapMap Phase 2 (Release 21) as reference panel. SNPs with imputation quality score (rsq) $\geq 0.5$ and minor allele frequency $\geq 1 \%$ were kept for analysis. Association analysis between asthma and individual SNPs was performed using a logistic regression model that
included allele dosage for each SNP and principal components to account for population structure. In family data, a robust sandwich estimation of the variance and a family cluster were used to take into account familial dependencies. Moreover, due to the complex sampling design of the GABRIELA study, survey regression techniques were used for this study to estimate robust standard errors ('svy' command in Stata). All analyses were performed using Stata ${ }^{\circledR}$ version 10 (distributed by Stata Corporation, College Station, Texas, USA).

## NTR: The Netherlands Twin Register

Subjects: NTR ${ }^{108,109}$ participants are ascertained because of the presence of twins or triplets in the family and consist of multiples, their parents, siblings and spouses. Twins are born in all strata of society and NTR represents a general sample from the Dutch population. In this study, 2,867 individuals were included (451 cases and 2,416 controls) with asthma definition: "ever doctor diagnosed asthma ignoring age of onset". Genotyping and QC: [La table sup 2 indique "Illumina 660K, Illumina Omni Express 1 mil, Affymetrix Perlegen and Affymetrix 6.0"] Genotype data were checked for European ancestry, Mendelian errors, gender inconsistencies and high genome-wide homozygosity. Checks for relatedness were carried out and only unrelated individuals were used in the analysis. Genotype data were further checked based on HardyWeinberg Equilibrium ( $1 \times 10^{-6} \mathrm{HWE}$ ), minor allele frequencies (MAF 0.01), SNP call rate ( $95 \%$ threshold of subjects with missing genotypes per SNP) and sample call rate ( $75 \%$ threshold of missing SNPs per subject).

Data Imputation and Analysis: The reference set used for imputation is HapMap phase II CEU data NCBI build 36 (UCSC hg18). Imputation was carried out using IMPUTE. Genome-wide association analysis was conducted using logistic regression (under an additive model) and including sex, age at examination and 4 principal components as covariates. Association analysis was conducted in PLINK, taking the uncertainty of the imputed genotypes into account.

## Rotterdam Study (RS-I, RS-II, RS-III)

Subjects: The Rotterdam Study is a prospective population based cohort study ongoing since 1990 in the city of Rotterdam in The Netherlands. Study design and objectives have been described elsewhere ${ }^{110}$. The first cohort (RS I) consists of 7,983 participants, aged 55 years and over. The second cohort (RS II) was recruited in 2000 with the same inclusion criteria and consists of 3,011 participants. The third cohort (RS IIII) consists of 3,932 participants, aged 45 years and over and was recruited in 2006. As of 2008, 14,926 subjects aged 45 years or over comprise the Rotterdam Study cohort. Asthma cases were collected by chart review. Cases were ascertained by 1) Doctor's diagnosis of Asthma and 2) no conflicting respiratory
diagnosis. Additionally, prescriptions for respiratory medicines were used for case finding. The number of cases and controls by study is shown in the Supplementary Table 1.

Genotyping and QC: Genotyping was done using the following arrays: Illumina Infinium II HumanHap550 chip v3.0 array (RS1), the HumanHap550 Duo Arrays and the Illumina Human610-Quad Arrays (RS-II), and the Human 610 Quad Arrays Illumina (RS-III). Exclusions were based on a call rate $<98 \%$, Hardy-Weinberg p-value $<10^{-6}$ and MAF $<0.01 \%$. SNP exclusions were based on a call rate $<98 \%$, Hardy-Weinberg p-value $<10^{-6}$ and MAF $<0.01 \%$.

Data Imputation and Analysis: We used the Markov Chain Haplotyping (MACH) package version 1.0.15 software (Rotterdam, The Netherlands; imputed to plus strand of NCBI build 36, HapMap release \#22) for the analyses of RSI and RSII. RSIII was imputed using MACH 1.0.16. For the analyses we used a MACH2DAT implemented in GRIMP, a high-speed pipeline for Genome Wide Association analyses ${ }^{34}$.

## DAGC Dutch Asthma Genetics Consortium

Subjects : The Dutch Asthma GWAS (DAG) cohort consists of in total 915 asthma cases and 986 controls, all from the northern of the Netherlands. The DAG cohort was genotyped in two phases and meta-analysed afterwards. For the first phase, 463 cases were selected from a trio and family study. The 469 controls were non-asthmatic spouses or pseudo-controls of untransmitted alleles in our trio design (GWAS I). For the second phase (GWAS II), 452 asthmatics were selected from previous clinical and genetic studies performed by our research institute. The 517 controls were selected from the COPACETIC study, a geographically matched population-based study on lung cancer screening in male smokers. All asthmatics had a physician's diagnosis of asthma, asthma symptoms, and BHR to either histamine or methacholine. BHR was measured with a metacholine or histamine challenge test, and defined as PC20 histamine. Controls had no asthma or COPD, nor any evidence of significant airway obstruction. ${ }^{111}$

Genotyping: Genotyping was performed on two platforms, 1. the Illumina HAPMAP 317K platform and 2. HAPMAP Illumina 370 Duo Chip. Quality control was applied; subjects were removed for analysis if they were not of Caucasian descent, had a low genotyping call rate (<95\%) or were discrepant or ambiguous for genetic sex. SNPs were deleted if the call rates were low (95\%), if they were not in Hardy-Weinberg Equilibrium ( $p<10 \mathrm{E}-04$ ), or if the minor allele frequency was $<0.05$.

Data Imputation and Analysis: Imputation was performed using BEAGLE and the HapMap2 CEU reference panel. All statistical analyses have been performed using PLINK v1.07.

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## Rotterdam Study (RS-I, RS-II, RS-III)

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## Supplementary References

1. Ferreira, M.A. et al. Identification of IL6R and chromosome 11q13.5 as risk loci for asthma. Lancet 378, 1006-14 (2011).
2. Howie, B.N., Donnelly, P. \& Marchini, J. A flexible and accurate genotype imputation method for the next generation of genome-wide association studies. PLoS Genet 5, e1000529 (2009).
3. Purcell, S. et al. PLINK: a tool set for whole-genome association and population-based linkage analyses. Am J Hum Genet 81, 559-75 (2007).
4. Chan-Yeung, M. et al. The Canadian Childhood Asthma Primary Prevention Study: outcomes at 7 years of age. J Allergy Clin Immunol 116, 49-55 (2005).
5. Kozyrskyj, A.L. et al. A novel study design to investigate the early-life origins of asthma in children (SAGE study). Allergy 64, 1185-93 (2009).
6. Moffatt, M.F. et al. A large-scale, consortium-based genomewide association study of asthma. $N$ Engl J Med 363, 1211-21 (2010).
7. Laprise, C. The Saguenay-Lac-Saint-Jean asthma familial collection: the genetics of asthma in a young founder population. Genes Immun 15, 247-55 (2014).
8. Splansky, G.L. et al. The Third Generation Cohort of the National Heart, Lung, and Blood Institute's Framingham Heart Study: design, recruitment, and initial examination. Am J Epidemiol 165, 132835 (2007).
9. Canoy, D. et al. Early growth and adult respiratory function in men and women followed from the fetal period to adulthood. Thorax 62, 396-402 (2007).
10. Day, N. et al. EPIC-Norfolk: study design and characteristics of the cohort. European Prospective Investigation of Cancer. Br J Cancer 80 Suppl 1, 95-103 (1999).
11. Vartiainen, E. et al. Thirty-five-year trends in cardiovascular risk factors in Finland. Int J Epidemiol 39, 504-18 (2010).
12. Aromaa, A. \& Koskinen, S. (eds.). Health and functional capacity in Finland. Baseline Results of the Health 2000 Health Examination Survey, (National Public Health Institute KTL, Helsinki, 2004).
13. Barker, D.J., Osmond, C., Forsen, T.J., Kajantie, E. \& Eriksson, J.G. Trajectories of growth among children who have coronary events as adults. N Engl J Med 353, 1802-9 (2005).
14. Raitakari, O.T. et al. Cohort profile: the cardiovascular risk in Young Finns Study. Int J Epidemiol 37, 1220-6 (2008).
15. Kantor, D.B. et al. Replication and fine mapping of asthma-associated loci in individuals of African ancestry. Hum Genet 132, 1039-47 (2013).
16. Lettre, G. et al. Genome-wide association study of coronary heart disease and its risk factors in 8,090 African Americans: the NHLBI CARe Project. PLoS Genet 7, e1001300 (2011).
17. Bild, D.E. et al. Multi-ethnic study of atherosclerosis: objectives and design. Am J Epidemiol 156, 871-81 (2002).
18. Kaufman, J.D. et al. Prospective study of particulate air pollution exposures, subclinical atherosclerosis, and clinical cardiovascular disease: The Multi-Ethnic Study of Atherosclerosis and Air Pollution (MESA Air). Am J Epidemiol 176, 825-37 (2012).
19. Manichaikul, A. et al. Analysis of family- and population-based samples in cohort genome-wide association studies. Hum Genet 131, 275-87 (2012).
20. Soler Artigas, M. et al. Genome-wide association and large-scale follow up identifies 16 new loci influencing lung function. Nat Genet 43, 1082-90 (2011).
21. THE ARIC INVESTIGATORS. The Atherosclerosis Risk in Communities (ARIC) Study: design and objectives. The ARIC investigators. Am J Epidemiol 129, 687-702 (1989).
22. Guan, Y. \& Stephens, M. Practical issues in imputation-based association mapping. PLoS Genet 4, e1000279 (2008).
23. R Core Team. R: A language and environment for statistical computing. R Foundation for Statistical Computing, Vienna, Austria. (2014).
24. Gudbjartsson, D.F. et al. Sequence variants affecting eosinophil numbers associate with asthma and myocardial infarction. Nat Genet 41, 342-7 (2009).
25. Halapi, E. et al. A sequence variant on $17 q 21$ is associated with age at onset and severity of asthma. Eur J Hum Genet 18, 902-8 (2010).
26. Marchini, J., Howie, B., Myers, S., McVean, G. \& Donnelly, P. A new multipoint method for genomewide association studies by imputation of genotypes. Nat Genet 39, 906-13 (2007).
27. Bisgaard, H. The Copenhagen Prospective Study on Asthma in Childhood (COPSAC): design, rationale, and baseline data from a longitudinal birth cohort study. Ann Allergy Asthma Immunol 93, 381-9 (2004).
28. Bisgaard, H., Hermansen, M.N., Loland, L., Halkjaer, L.B. \& Buchvald, F. Intermittent inhaled corticosteroids in infants with episodic wheezing. N Engl J Med 354, 1998-2005 (2006).
29. Bisgaard, H. et al. Childhood asthma after bacterial colonization of the airway in neonates. $N$ Engl J Med 357, 1487-95 (2007).
30. Li, Y., Willer, C.J., Ding, J., Scheet, P. \& Abecasis, G.R. MaCH: using sequence and genotype data to estimate haplotypes and unobserved genotypes. Genet Epidemiol 34, 816-34 (2010).
31. Olsen, J. et al. The Danish National Birth Cohort--its background, structure and aim. Scand J Public Health 29, 300-7 (2001).
32. Ryckman, K.K. et al. Replication of a genome-wide association study of birth weight in preterm neonates. J Pediatr 160, 19-24 e4 (2012).
33. Jaddoe, V.W. et al. The Generation R Study: design and cohort update 2012. Eur J Epidemiol 27, 739-56 (2012).
34. Estrada, K. et al. GRIMP: a web- and grid-based tool for high-speed analysis of large-scale genomewide association using imputed data. Bioinformatics 25, 2750-2 (2009).
35. Heinrich, J. et al. Allergens and endotoxin on mothers' mattresses and total immunoglobulin E in cord blood of neonates. Eur Respir J 20, 617-23 (2002).
36. Heinrich, J. et al. [Two German Birth Cohorts: GINIplus and LISAplus]. Bundesgesundheitsblatt Gesundheitsforschung Gesundheitsschutz 55, 864-74 (2012).
37. Berg, A. et al. Impact of early feeding on childhood eczema: development after nutritional intervention compared with the natural course - the GINIplus study up to the age of 6 years. Clin Exp Allergy 40, 627-36 (2010).
38. Custovic, A., Simpson, B.M., Murray, C.S., Lowe, L. \& Woodcock, A. The National Asthma Campaign Manchester Asthma and Allergy Study. Pediatr Allergy Immunol 13 Suppl 15, 32-7 (2002).
39. Lowe, L.A. et al. Wheeze phenotypes and lung function in preschool children. Am J Respir Crit Care Med 171, 231-7 (2005).
40. Murray, C.S. et al. Lung function at one month of age as a risk factor for infant respiratory symptoms in a high risk population. Thorax 57, 388-92 (2002).
41. Nicolaou, N.C. et al. Exhaled breath condensate pH and childhood asthma: unselected birth cohort study. Am J Respir Crit Care Med 174, 254-9 (2006).
42. Nicolaou, N.C. et al. Day-care attendance, position in sibship, and early childhood wheezing: a population-based birth cohort study. J Allergy Clin Immunol 122, 500-6 e5 (2008).
43. Newnham, J.P., Evans, S.F., Michael, C.A., Stanley, F.J. \& Landau, L.I. Effects of frequent ultrasound during pregnancy: a randomised controlled trial. Lancet 342, 887-91 (1993).
44. Wellcome Trust Case Control, C. Genome-wide association study of 14,000 cases of seven common diseases and 3,000 shared controls. Nature 447, 661-78 (2007).
45. Anderson, H.R., Pottier, A.C. \& Strachan, D.P. Asthma from birth to age 23: incidence and relation to prior and concurrent atopic disease. Thorax 47, 537-42 (1992).
46. Strachan, D.P., Butland, B.K. \& Anderson, H.R. Incidence and prognosis of asthma and wheezing illness from early childhood to age 33 in a national British cohort. BMJ 312, 1195-9 (1996).
47. Butland, B.K. \& Strachan, D.P. Asthma onset and relapse in adult life: the British 1958 birth cohort study. Ann Allergy Asthma Immunol 98, 337-43 (2007).
48. Strachan, D.P. et al. Lifecourse influences on health among British adults: effects of region of residence in childhood and adulthood. Int J Epidemiol 36, 522-31 (2007).
49. Wellcome Trust Case Control Consortium. Genome-wide association study of 14,000 cases of seven common diseases and 3,000 shared controls. Nature 447, 661-78 (2007).
50. Barrett, J.C. et al. Genome-wide association study and meta-analysis find that over 40 loci affect risk of type 1 diabetes. Nat Genet 41, 703-7 (2009).
51. Aulchenko, Y.S., Struchalin, M.V. \& van Duijn, C.M. ProbABEL package for genome-wide association analysis of imputed data. BMC Bioinformatics 11, 134 (2010).
52. Moore, W.C. et al. Characterization of the severe asthma phenotype by the National Heart, Lung, and Blood Institute's Severe Asthma Research Program. J Allergy Clin Immunol 119, 405-13 (2007).
53. Moore, W.C. et al. Identification of asthma phenotypes using cluster analysis in the Severe Asthma Research Program. Am J Respir Crit Care Med 181, 315-23 (2010).
54. Sankararaman, S., Sridhar, S., Kimmel, G. \& Halperin, E. Estimating local ancestry in admixed populations. Am J Hum Genet 82, 290-303 (2008).
55. Price, A.L. et al. Principal components analysis corrects for stratification in genome-wide association studies. Nat Genet 38, 904-9 (2006).
56. Pritchard, J.K., Stephens, M. \& Donnelly, P. Inference of population structure using multilocus genotype data. Genetics 155, 945-59 (2000).
57. American Thoracic Society. Standards for the diagnosis and care of patients with chronic obstructive pulmonary disease (COPD) and asthma. This official statement of the American Thoracic Society was adopted by the ATS Board of Directors, November 1986. Am Rev Respir Dis 136, 225-44 (1987).
58. The International Study of Asthma and Allergies in Childhood (ISAAC) Steering Committee. Worldwide variations in the prevalence of asthma symptoms: the International Study of Asthma and Allergies in Childhood (ISAAC). Eur Respir J 12, 315-35 (1998).
59. Bonilla, C. et al. The 8818 G allele of the agouti signaling protein (ASIP) gene is ancestral and is associated with darker skin color in African Americans. Hum Genet 116, 402-6 (2005).
60. Mathias, R.A. et al. A genome-wide association study on African-ancestry populations for asthma. $J$ Allergy Clin Immunol 125, 336-346 e4 (2010).
61. Epstein, M.P., Duren, W.L. \& Boehnke, M. Improved inference of relationship for pairs of individuals. Am J Hum Genet 67, 1219-31 (2000).
62. Wu, B., Liu, N. \& Zhao, H. PSMIX: an R package for population structure inference via maximum likelihood method. BMC Bioinformatics 7, 317 (2006).
63. Hirota, T. et al. Genome-wide association study identifies three new susceptibility loci for adult asthma in the Japanese population. Nat Genet 43, 893-6 (2011).
64. Noguchi, E. et al. Genome-wide association study identifies HLA-DP as a susceptibility gene for pediatric asthma in Asian populations. PLoS Genet 7, e1002170 (2011).
65. Kauffmann, F. et al. Epidemiological study of the genetics and environment of asthma, bronchial hyperresponsiveness, and atopy: phenotype issues. Am J Respir Crit Care Med 156, S123-9 (1997).
66. Kauffmann, F. et al. EGEA (Epidemiological study on the Genetics and Environment of Asthma, bronchial hyperresponsiveness and atopy)-- descriptive characteristics. Clin Exp Allergy 29 Suppl 4, 17-21 (1999).
67. Boyd, A. et al. Cohort Profile: the 'children of the 90 s'--the index offspring of the Avon Longitudinal Study of Parents and Children. Int J Epidemiol 42, 111-27 (2013).
68. ECRHS. The European Community Respiratory Health Survey II. Eur Respir J 20, 1071-9 (2002).
69. Burney, P.G., Luczynska, C., Chinn, S. \& Jarvis, D. The European Community Respiratory Health Survey. Eur Respir J 7, 954-60 (1994).
70. Wickman, M., Kull, I., Pershagen, G. \& Nordvall, S.L. The BAMSE project: presentation of a prospective longitudinal birth cohort study. Pediatr Allergy Immunol 13 Suppl 15, 11-3 (2002).
71. Melen, E. et al. Interactions between glutathione S-transferase P1, tumor necrosis factor, and traffic-related air pollution for development of childhood allergic disease. Environ Health Perspect 116, 1077-84 (2008).
72. James, A.L. et al. Changes in the prevalence of asthma in adults since 1966: the Busselton health study. Eur Respir J 35, 273-8 (2010).
73. Genuneit, J. et al. The GABRIEL Advanced Surveys: study design, participation and evaluation of bias. Paediatr Perinat Epidemiol 25, 436-47 (2011).
74. Polonikov, A.V. et al. Promoter polymorphism G-50T of a human CYP2J2 epoxygenase gene is associated with common susceptibility to asthma. Chest 132, 120-6 (2007).
75. Polonikov, A.V. et al. The relationship between polymorphisms in the glutamate cysteine ligase gene and asthma susceptibility. Respir Med 101, 2422-4 (2007).
76. British guideline on the management of asthma. Thorax 58 Suppl 1, i1-94 (2003).
77. Moffatt, M.F. et al. Genetic variants regulating ORMDL3 expression contribute to the risk of childhood asthma. Nature 448, 470-3 (2007).
78. Weiland, S.K. et al. Prevalence of respiratory and atopic disorders among children in the East and West of Germany five years after unification. Eur Respir J 14, 862-70 (1999).
79. Weiland, S.K. et al. Phase II of the International Study of Asthma and Allergies in Childhood (ISAAC II): rationale and methods. Eur Respir J 24, 406-12 (2004).
80. Lau, S. et al. The development of childhood asthma: lessons from the German Multicentre Allergy Study (MAS). Paediatr Respir Rev 3, 265-72 (2002).
81. Nickel, R. et al. How should a birth cohort study be organised? Experience from the German MAS cohort study. Paediatr Respir Rev 3, 169-76 (2002).
82. Zuidgeest, M.G. et al. Persistence of asthma medication use in preschool children. Respir Med 102, 1446-51 (2008).
83. Wijga, A.H. et al. Cohort profile: the prevention and incidence of asthma and mite allergy (PIAMA) birth cohort. Int J Epidemiol 43, 527-35 (2014).
84. Brunekreef, B. et al. The prevention and incidence of asthma and mite allergy (PIAMA) birth cohort study: design and first results. Pediatr Allergy Immunol 13 Suppl 15, 55-60 (2002).
85. Downs, S.H. et al. Reduced exposure to PM10 and attenuated age-related decline in lung function. N Engl J Med 357, 2338-47 (2007).
86. Castro-Giner, F. et al. TNFA -308G>A in two international population-based cohorts and risk of asthma. Eur Respir J 32, 350-61 (2008).
87. Imboden, M. et al. Decreased PM10 exposure attenuates age-related lung function decline: genetic variants in p53, p21, and CCND1 modify this effect. Environ Health Perspect 117, 1420-7 (2009).
88. Freidin, M.B., Puzyrev, V.P., Ogorodova, L.M., Kobiakova, O.S. \& Kulmanakova, I.M. [Polymorphism of interleukins and interleukin receptor genes: population distribution and association with atopic bronchial asthma]. Genetika 38, 1710-8 (2002).
89. Freidin, M.B., Kobyakova, O.S., Ogorodova, L.M. \& Puzyrev, V.P. Association of polymorphisms in the human IL4 and IL5 genes with atopic bronchial asthma and severity of the disease. Comp Funct Genomics 4, 346-50 (2003).
90. Sigsgaard, T., Hjort, C., Omland, $\varnothing$., Miller, M.R. \& Pedersen, O.F. Respiratory Health and Allergy Among Young Farmers and Non-Farming Rural Males in Denmark. Journal of Agromedicine 4, 6378 (1997).
91. Omland, O., Sigsgaard, T., Hjort, C., Pedersen, O.F. \& Miller, M.R. Lung status in young Danish rurals: the effect of farming exposure on asthma-like symptoms and lung function. Eur Respir J 13, 31-7 (1999).
92. Smit, L.A. et al. Exposure-response analysis of allergy and respiratory symptoms in endotoxinexposed adults. Eur Respir J 31, 1241-8 (2008).
93. Smit, L.A. et al. Ex vivo cytokine release reflects sensitivity to occupational endotoxin exposure. Eur Respir J 34, 795-802 (2009).
94. Samadi, S. et al. Allergy among veterinary medicine students in The Netherlands. Occup Environ Med 69, 48-55 (2012).
95. Jacobs, J.H., Meijster, T., Meijer, E., Suarthana, E. \& Heederik, D. Wheat allergen exposure and the prevalence of work-related sensitization and allergy in bakery workers. Allergy 63, 1597-604 (2008).
96. Schlunssen, V., Schaumburg, I., Taudorf, E., Mikkelsen, A.B. \& Sigsgaard, T. Respiratory symptoms and lung function among Danish woodworkers. J Occup Environ Med 44, 82-98 (2002).
97. Jacobsen, G., Schlunssen, V., Schaumburg, I., Taudorf, E. \& Sigsgaard, T. Longitudinal lung function decline and wood dust exposure in the furniture industry. Eur Respir J 31, 334-42 (2008).
98. Pronk, A. et al. Respiratory symptoms, sensitization, and exposure response relationships in spray painters exposed to isocyanates. Am J Respir Crit Care Med 176, 1090-7 (2007).
99. Pronk, A. et al. Different respiratory phenotypes are associated with isocyanate exposure in spray painters. Eur Respir J 33, 494-501 (2009).
100. Basinas, l. et al. Sensitisation to common allergens and respiratory symptoms in endotoxin exposed workers: a pooled analysis. Occup Environ Med 69, 99-106 (2012).
101. Haldar, P. et al. Cluster analysis and clinical asthma phenotypes. Am J Respir Crit Care Med 178, 218-24 (2008).
102. Haldar, P. et al. Mepolizumab and exacerbations of refractory eosinophilic asthma. N Engl J Med 360, 973-84 (2009).
103. Proceedings of the ATS workshop on refractory asthma: current understanding, recommendations, and unanswered questions. American Thoracic Society. Am J Respir Crit Care Med 162, 2341-51 (2000).
104. Chung, K.F. et al. Difficult/therapy-resistant asthma: the need for an integrated approach to define clinical phenotypes, evaluate risk factors, understand pathophysiology and find novel therapies. ERS Task Force on Difficult/Therapy-Resistant Asthma. European Respiratory Society. Eur Respir J 13, 1198-208 (1999).
105. Ayres, J.G., Miles, J.F. \& Barnes, P.J. Brittle asthma. Thorax 53, 315-21 (1998).
106. Okaneya, T., Mizusawa, H., Taguchi, I. \& Yoneyama, T. [Clinical evaluation of tandem PSA for the diagnosis and follow-up of prostate cancer]. Hinyokika Kiyo 42, 861-7 (1996).
107. Stanojevic, S. et al. Reference ranges for spirometry across all ages: a new approach. Am J Respir Crit Care Med 177, 253-60 (2008).
108. Boomsma, D.I. et al. Genome-wide association of major depression: description of samples for the GAIN Major Depressive Disorder Study: NTR and NESDA biobank projects. Eur J Hum Genet 16, 335-42 (2008).
109. Willemsen, G. et al. The Netherlands Twin Register biobank: a resource for genetic epidemiological studies. Twin Res Hum Genet 13, 231-45 (2010).
110. Hofman, A. et al. The Rotterdam Study: 2012 objectives and design update. Eur J Epidemiol 26, 657-86 (2011).
111. Nieuwenhuis, M.A. et al. Combining genomewide association study and lung eQTL analysis provides evidence for novel genes associated with asthma. Allergy (2016).

[^0]:    ${ }^{\text {a }}$ EAF $=$ Effect allele frequency
    ${ }^{\text {b }}$ Odds-ratios (ORs) and 95\% Confidence Intervals (CI) were computed for the effect allele (random \& fixed-effects models)
    ${ }^{\text {c }}$ random is P -value for test of association between SNP and asthma under a random-effects model
    ${ }^{d}{ }^{\text {fixed }}$ is P -value for test of association between SNP and asthma under a fixed-effects model
    ${ }^{\text {eP }}{ }_{\text {het }}$ is the P -value for test of heterogeneity across studies with the use of Cochran's test

[^1]:    ${ }^{\text {a }}$ Cytogenetic position of the chromosomal region harboring the lead SNP
    ${ }^{\mathrm{b}}$ Genes coding for proteins; the gene where eventually the lead SNP lies is first indicated followed by the previous gene and next gene
    ${ }^{\text {cR }}$ =reference allele / $\mathrm{E}=$ effect allele.
    ${ }^{\text {d O Odds-ratios (ORs) and } 95 \% \text { Confidence Intervals (CI) were computed for the effect allele under a random-effects model }}$
    ${ }^{e} P_{\text {random }}$ is the $P$-value for test of association between SNP and asthma under a random-effects model
    ${ }^{f} P_{\text {fixed }}$ is the $P$-value for test of association between SNP and asthma under a fixed-effects model
    ${ }^{8} P_{\text {het }}$ is the P-value for test of heterogeneity across studies with the use of Cochran's test

[^2]:    ${ }^{2} \mathrm{R}=$ reference allele / $\mathrm{E}=$ effect allele
    ${ }^{\text {b }}$ Odds-ratios (ORs) and $95 \%$ Confidence Intervals (CI) were computed for the effect allele under a random-effects model
    ${ }^{\mathrm{CP}}$ random for test of association between SNP and asthma under a random-effects model
    ${ }^{d} P_{\text {fixed }}$ for test of association between SNP and asthma under a fixed-effects model
    ${ }^{e} P_{\text {het }}$ for test of heterogeneity across studies with the use of Cochran's $Q$ test

