**Supplementary Table 1. Results of Mendelian randomization analyses and sensitivity analyses for smoking initiation on schizophrenia**

|  |  |  |  |
| --- | --- | --- | --- |
| **Method** | **ORa** | **95% CIb** | ***P*-value** |
| Simple median | 0.263 | 0.042 - 1.660 | 0.156 |
| Weighted median | 0.261 | 0.043 - 1.560 | 0.141 |
| IVW | 0.318 | 0.073 - 1.394 | 0.129 |
| MR-Egger | 12.718 | 0.021 - >20.0 | 0.438 |
| intercept | 0.969 | 0.921 - 1.021 | 0.248 |

a OR: Odds ratio.

b 95% CI: 95% Confidence interval.

**Supplementary Table 2. The associations of individual SNPs (used as MR instruments) with smoking initiation and schizophrenia**

|  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- |
| **Smoking initiation-SNPa** | **Chrb** | **Position (bp)** | **Allelesc** | **Smoking initiation results** | | **Schizophrenia results** | |
| ***P*-value** | **βd** | ***P*-value** | **βd** |
| rs209584 | 1 | 40852821 | A/G | 7.89E-06 | 0.014 | 0.117 | -0.090 |
| rs11205679 | 1 | 50541643 | G/C | 1.78E-06 | -0.007 | 0.051 | 0.043 |
| rs3856101 | 1 | 182022938 | A/G | 4.04E-07 | -0.008 | 0.327 | -0.025 |
| rs939584 | 2 | 621558 | T/C | 2.89E-06 | 0.011 | 0.689 | -0.015 |
| rs10168190 | 2 | 59983904 | A/C | 8.97E-06 | -0.007 | 0.297 | -0.024 |
| rs73934251 | 2 | 62727986 | C/A | 5.07E-07 | 0.008 | 0.122 | -0.037 |
| rs3770582 | 2 | 169819824 | G/A | 6.19E-06 | -0.007 | 0.823 | 0.007 |
| rs6436573 | 2 | 226459822 | G/C | 1.26E-06 | -0.009 | 0.004 | 0.029 |
| rs1814463 | 3 | 150487902 | G/A | 1.19E-07 | -0.008 | 0.610 | 0.011 |
| rs10212571 | 3 | 161878203 | G/C | 8.25E-06 | -0.007 | 0.010 | 0.074 |
| rs77068703 | 4 | 19574610 | T/C | 5.95E-06 | 0.008 | 0.145 | 0.038 |
| rs12331784 | 4 | 140912403 | A/G | 3.26E-06 | 0.007 | 0.069 | 0.042 |
| rs10517796 | 4 | 164910589 | C/T | 6.80E-06 | 0.007 | 0.564 | -0.013 |
| rs2680793 | 5 | 23030394 | T/C | 1.59E-07 | 0.008 | 0.890 | 0.003 |
| rs114656770 | 5 | 122936301 | C/T | 6.92E-06 | -0.012 | 0.318 | -0.035 |
| rs11153371 | 6 | 112894354 | C/T | 3.50E-06 | 0.007 | 0.197 | -0.029 |
| rs34072054 | 6 | 131446872 | C/A | 1.78E-06 | 0.009 | 0.677 | -0.016 |
| rs2030958 | 7 | 1076010 | G/T | 1.53E-06 | -0.009 | 0.132 | -0.062 |
| rs4729215 | 7 | 95473550 | A/G | 6.92E-06 | 0.007 | 0.425 | -0.018 |
| rs117036946 | 8 | 13287799 | A/G | 6.22E-08 | -0.023 | 0.670 | 0.029 |
| rs3924101 | 9 | 126187698 | C/T | 9.94E-06 | 0.006 | 0.114 | -0.053 |
| rs12379582 | 9 | 129164861 | C/T | 2.40E-06 | 0.009 | 0.639 | 0.018 |
| rs10733835 | 10 | 68971354 | A/C | 9.58E-06 | 0.007 | 0.037 | -0.047 |
| rs11030104 | 11 | 27684517 | A/G | 9.45E-06 | 0.006 | 0.392 | -0.018 |
| rs7118410 | 11 | 29403818 | G/A | 2.89E-06 | -0.016 | 0.113 | -0.070 |
| rs1860368 | 11 | 63994692 | C/T | 6.43E-06 | 0.007 | 0.542 | 0.019 |
| rs291217 | 11 | 86033229 | G/A | 6.80E-06 | 0.007 | 0.913 | 0.002 |
| rs28652571 | 16 | 61910001 | A/G | 6.80E-06 | -0.007 | 0.852 | -0.004 |
| rs643614 | 16 | 77137196 | G/A | 4.48E-06 | -0.009 | 0.414 | 0.024 |
| rs2156008 | 18 | 53281244 | A/G | 5.32E-07 | 0.008 | 0.196 | -0.029 |
| rs5752763 | 22 | 28999402 | A/G | 8.49E-07 | 0.008 | 0.748 | -0.011 |

a SNP: Single nucleotide polymorphism; b Chr: Chromosome; c Alleles: shown as other allele/effect allele; d β: effect sizes for the effect allele.

**Supplementary Table 3. Results of Mendelian randomization analyses and sensitivity analyses for smoking cessation on schizophrenia**

|  |  |  |  |
| --- | --- | --- | --- |
| **Method** | **ORa** | **95% CIb** | ***P*-value** |
| Simple median | 1.087 | 0.133 - 8.873 | 0.938 |
| Weighted median | 1.093 | 0.126 - 9.507 | 0.935 |
| IVW | 1.397 | 0.259 - 7.538 | 0.698 |
| MR-Egger | 0.474 | 0.000 - >10.0 | 0.907 |
| intercept | 1.016 | 0.845 - 1.221 | 0.865 |

a OR: Odds ratio.

b 95% CI: 95% Confidence interval.

**Supplementary Table 4. The associations of individual SNPs (used as MR instruments) with smoking cessation and schizophrenia.**

|  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- |
| **Smoking cessation-SNPa** | **Chrb** | **Position (bp)** | **Allelesc** | **Smoking cessation results** | | **Schizophrenia results** | |
| ***P*-value** | **βd** | ***P*-value** | **βd** |
| rs12041121 | 1 | 71946767 | A/G | 9.55E-06 | 0.015 | 0.174 | 0.041 |
| rs10925273 | 1 | 237087952 | C/A | 9.79E-06 | 0.014 | 0.468 | -0.028 |
| rs6852063 | 4 | 164962967 | T/G | 6.50E-06 | -0.019 | 0.966 | -0.002 |
| rs1374541 | 11 | 41452393 | C/T | 1.31E-07 | 0.014 | 0.967 | 0.001 |
| rs7934387 | 11 | 127338788 | G/A | 8.59E-07 | 0.012 | 0.422 | 0.023 |
| rs6019095 | 20 | 47000239 | A/C | 8.45E-06 | -0.013 | 0.284 | 0.045 |

a SNP: Single nucleotide polymorphism;

b Chr: Chromosome;

c Alleles: shown as other allele/effect allele;

d β: effect sizes for the effect allele.

**Supplementary Table 5. Results of Mendelian randomization analyses and sensitivity analyses for age of smoking initiation and schizophrenia**

|  |  |  |  |
| --- | --- | --- | --- |
| **Method** | **ORa** | **95% CIb** | ***P*-value** |
| Simple median | 0.689 | 0.333 - 1.425 | 0.314 |
| Weighted median | 0.785 | 0.399 - 1.547 | 0.483 |
| IVW | 0.976 | 0.589 - 1.619 | 0.927 |
| MR-Egger | 1.082 | 0.048 - >10.0 | 0.960 |
| (intercept) | 0.995 | 0.864 - 1.146 | 0.948 |

a OR: Odds ratio.

b 95% CI: 95% Confidence interval.

**Supplementary Table 6. The associations of individual SNPs (used as MR instruments) with age of smoking initiation and schizophrenia**

|  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- |
| **Age of smoking initiation-SNPa** | **Chrb** | **Position (bp)** | **Allelesc** | **Age of smoking initiation results** | | **Schizophrenia results** | |
| ***P*-value** | **βd** | ***P*-value** | **β**d |
| rs13107340 | 4 | 151116764 | G/T | 4.20E-06 | 0.040 | 0.480 | -0.016 |
| rs28722815 | 7 | 50654755 | G/A | 9.81E-06 | -0.075 | 0.436 | -0.053 |
| rs1014114 | 7 | 108041794 | G/T | 3.39E-06 | 0.047 | 0.281 | 0.033 |
| rs7043313 | 9 | 793989 | A/C | 2.12E-06 | -0.055 | 0.544 | 0.021 |
| rs12366735 | 12 | 85014953 | G/A | 8.56E-06 | 0.054 | 0.326 | -0.046 |
| rs4984585 | 15 | 96052500 | T/C | 3.97E-06 | -0.039 | 0.301 | -0.024 |
| rs4817902 | 21 | 39261724 | C/T | 1.77E-06 | 0.042 | 0.269 | -0.039 |

a SNP: Single nucleotide polymorphism.

b Chr: Chromosome.

c Alleles: shown as other allele/effect allele;

d β: effect sizes for the effect allele.