

Supplementary

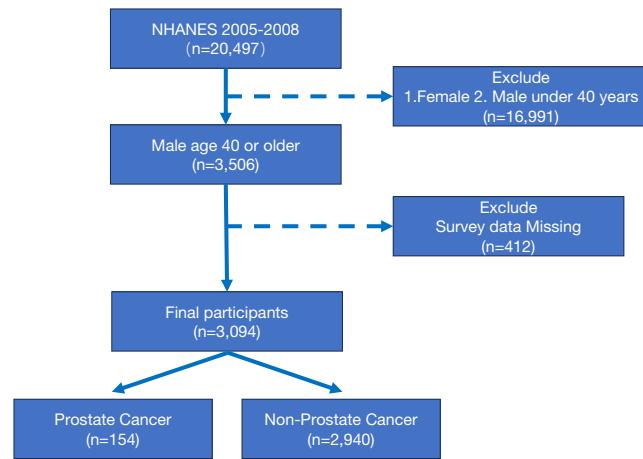


Figure S1 Flowchart of sample selection from NHANES (2005–2008). NHANES, National Health and Nutrition Examination Survey.

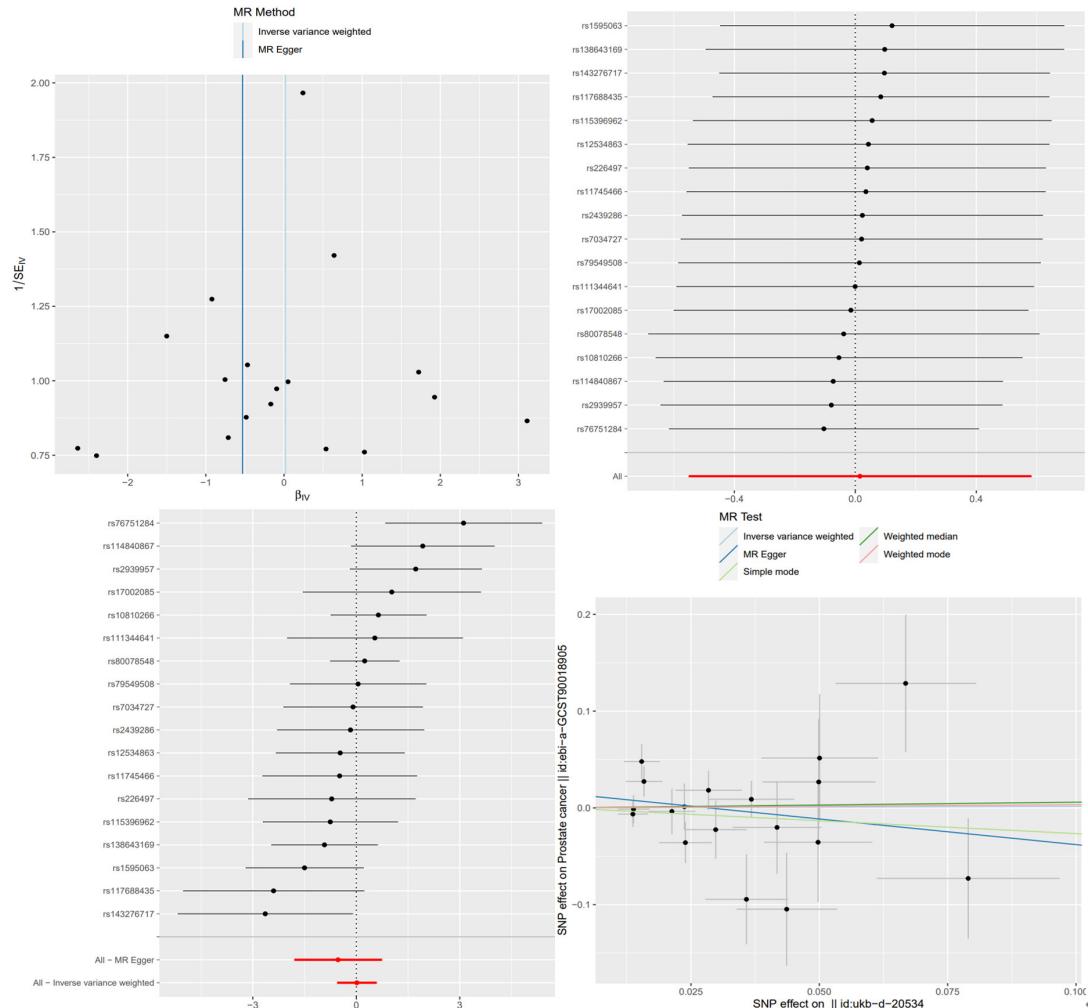


Figure S2 Forest plot, funnel plot, scatter and sensitivity analysis for Sleeping too much in prostate cancer (Mendelian Randomization, MR).

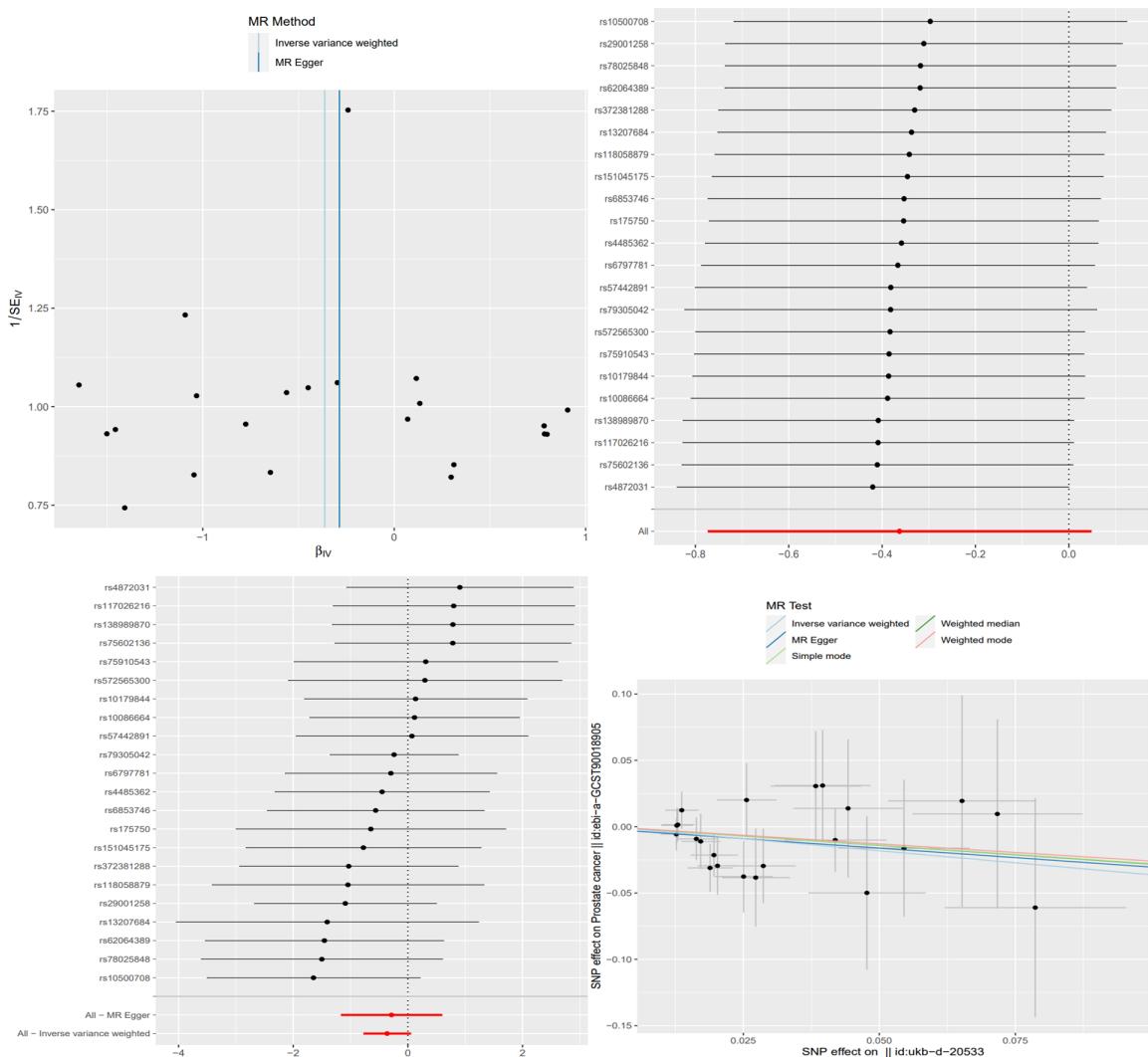


Figure S3 Forest plot, funnel plot, scatter and sensitivity analysis for Trouble falling asleep in prostate cancer (Mendelian Randomization, MR).

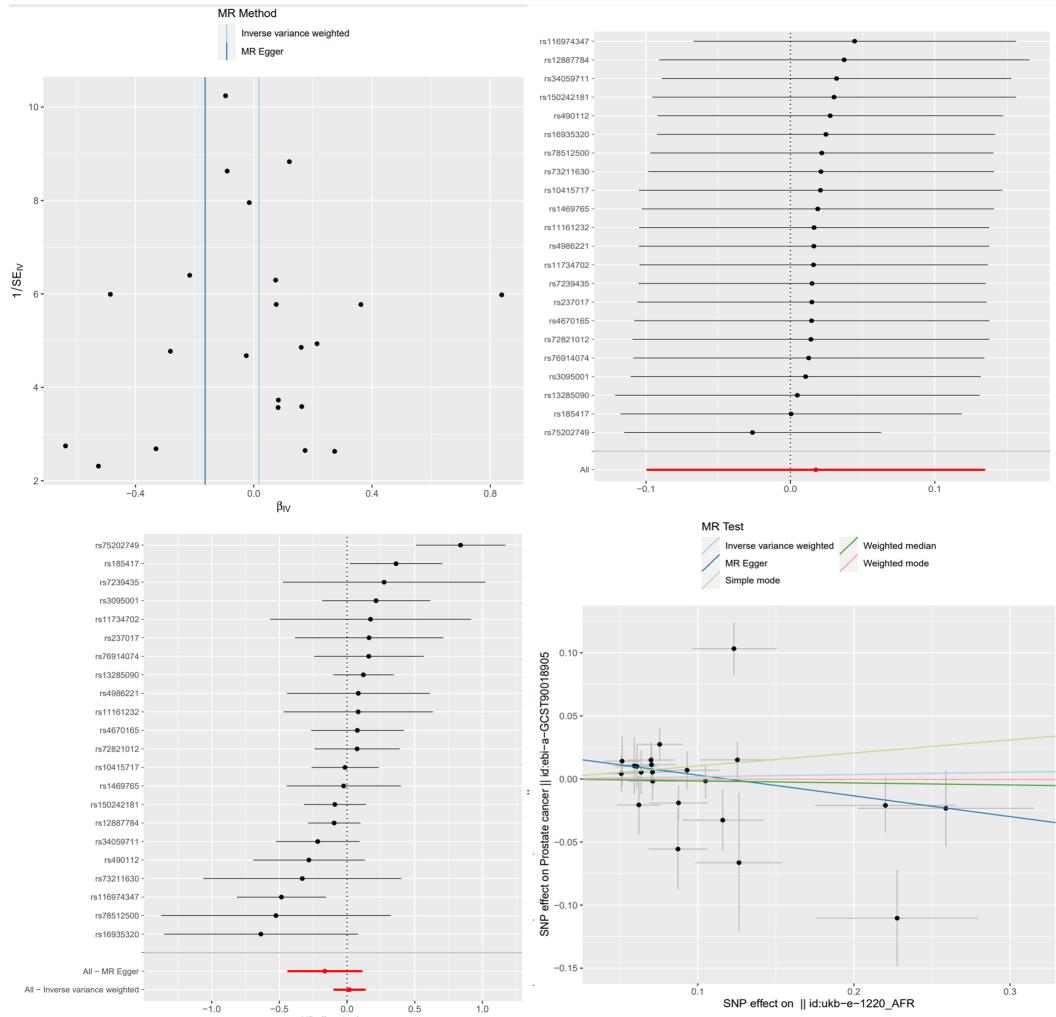


Figure S4 Forest plot, funnel plot, scatter and sensitivity analysis for Daytime dozing / sleeping (narcolepsy) in prostate cancer (Mendelian Randomization, MR).

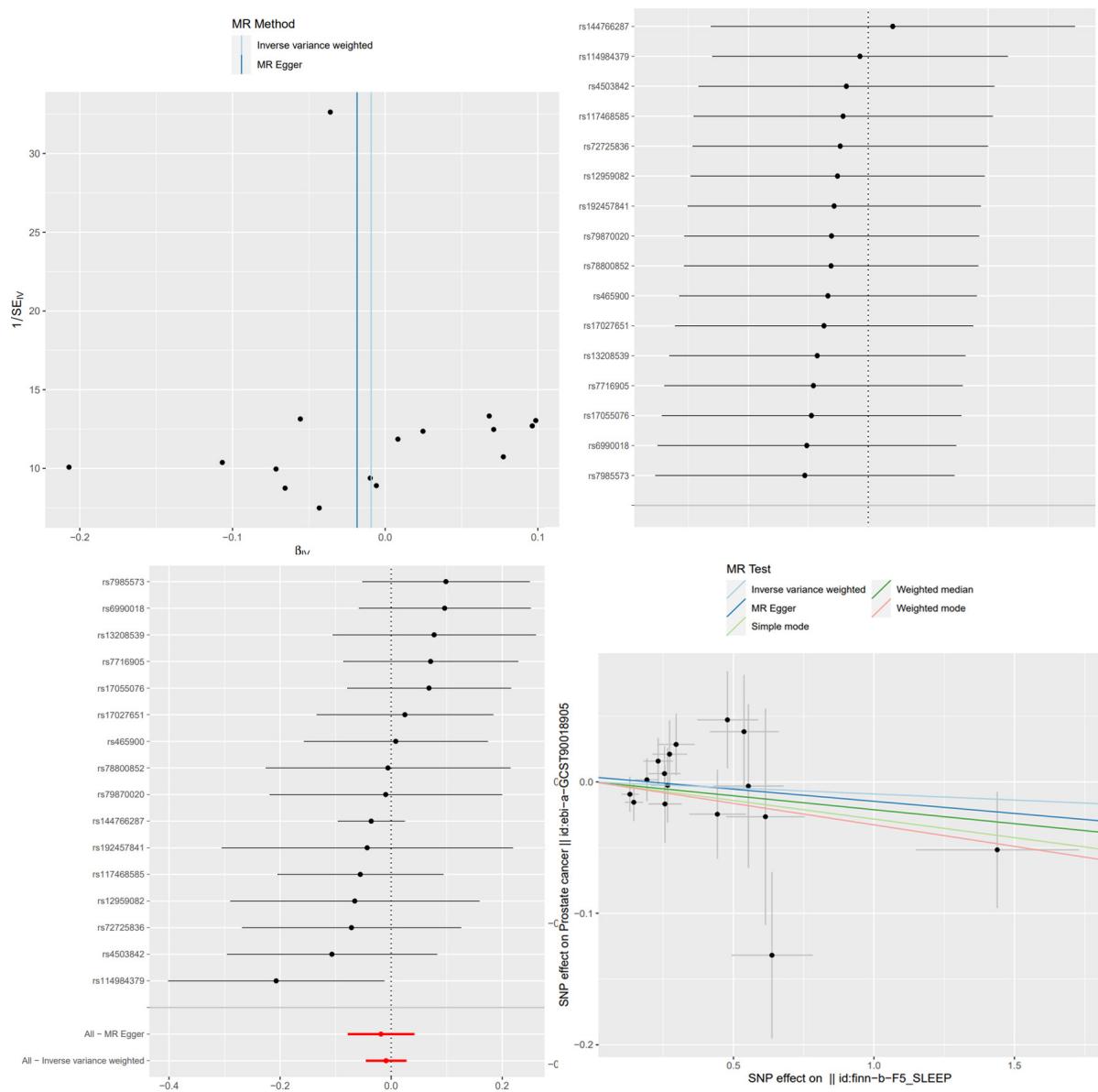


Figure S5 Forest plot, funnel plot, scatter and sensitivity analysis for sleep disorders in prostate cancer (Mendelian Randomization, MR).

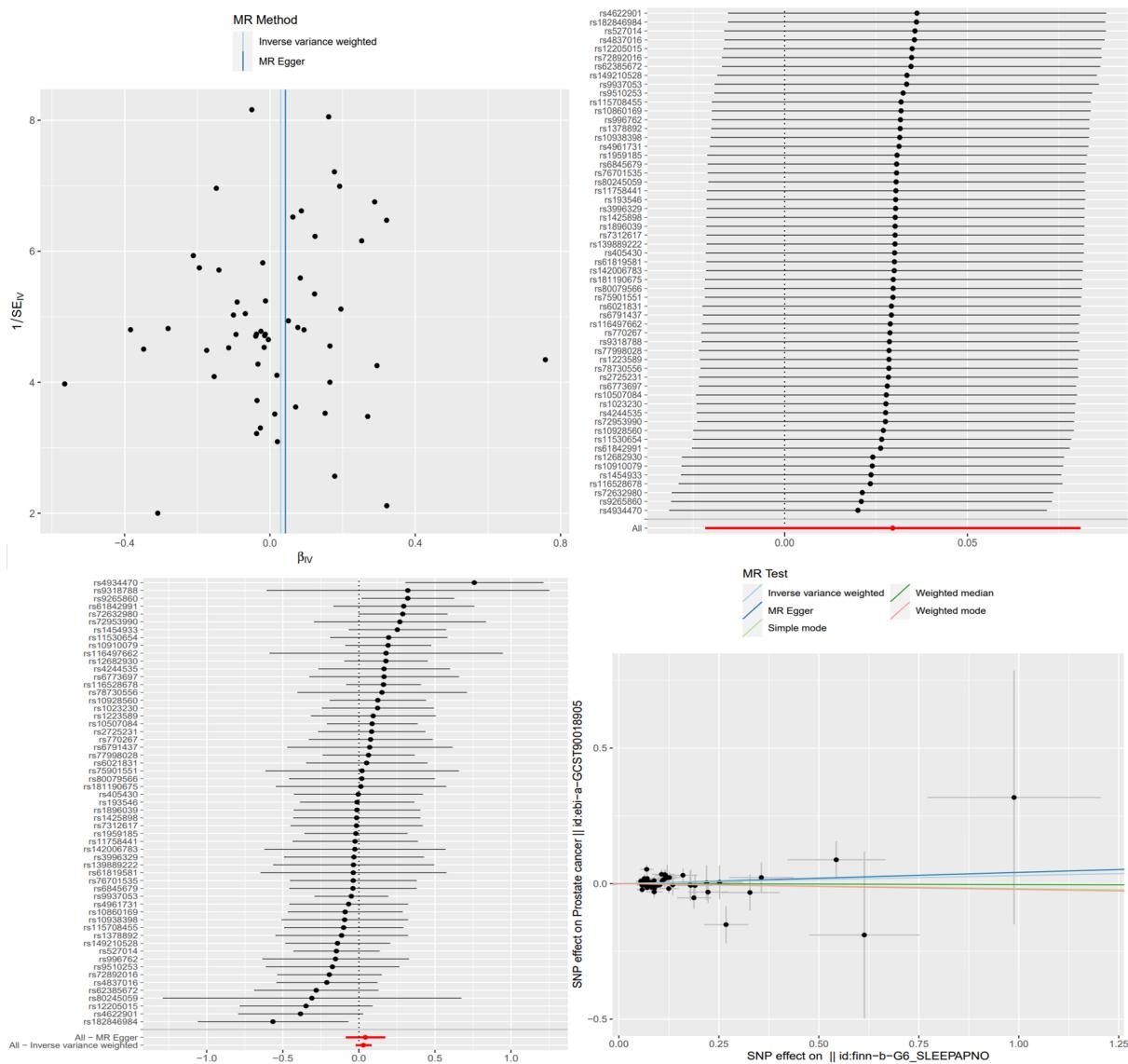


Figure S6 Forest plot, funnel plot, scatter and sensitivity analysis for Sleep apnea in prostate cancer (Mendelian Randomization, MR).

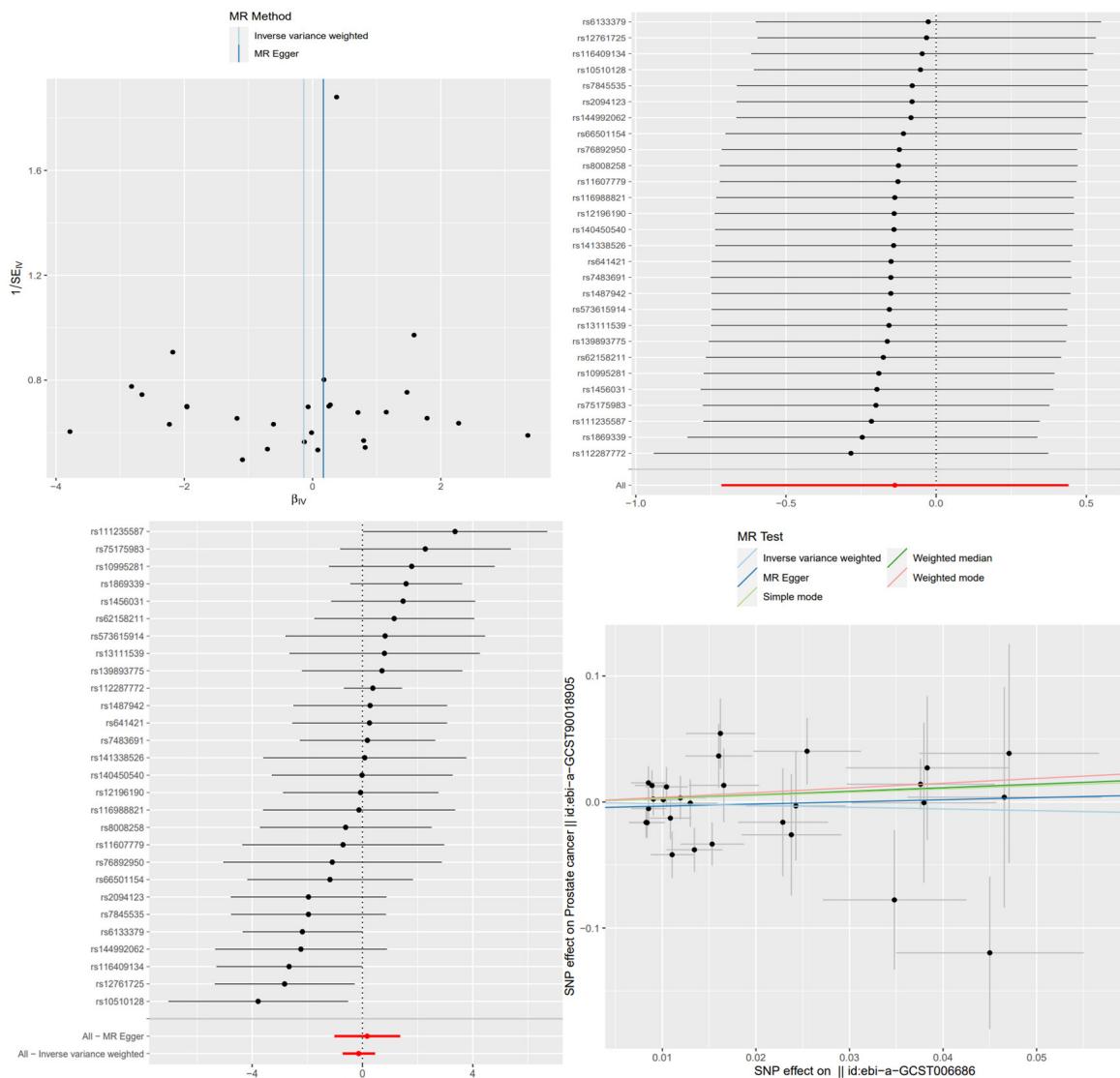


Figure S7 Forest plot, funnel plot, scatter and sensitivity analysis for Sleep duration (undersleepers) in prostate cancer (Mendelian Randomization, MR).

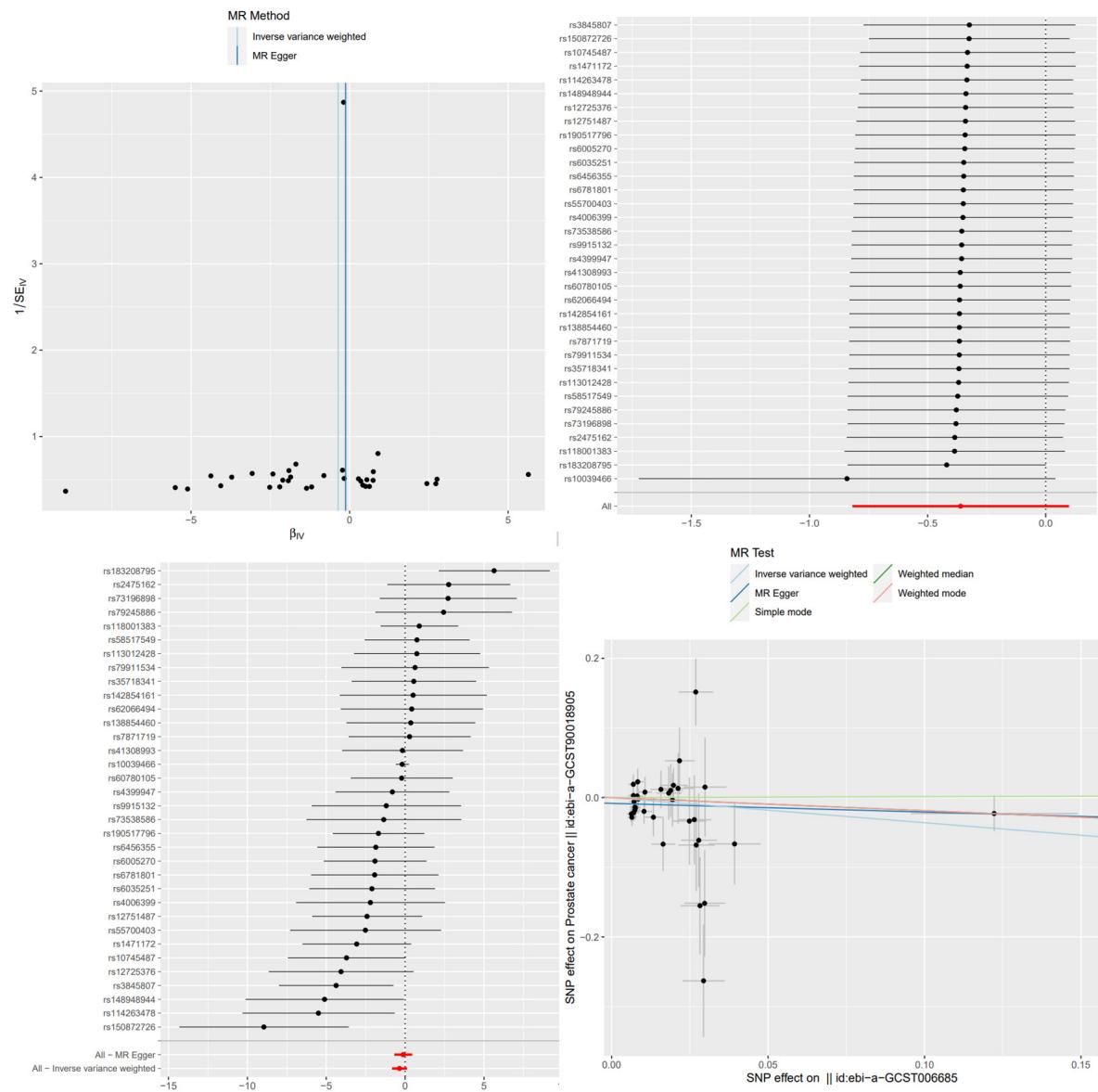


Figure S8 Forest plot, funnel plot, scatter and sensitivity analysis for Sleep duration (oversleepers) in prostate cancer (Mendelian Randomization, MR).

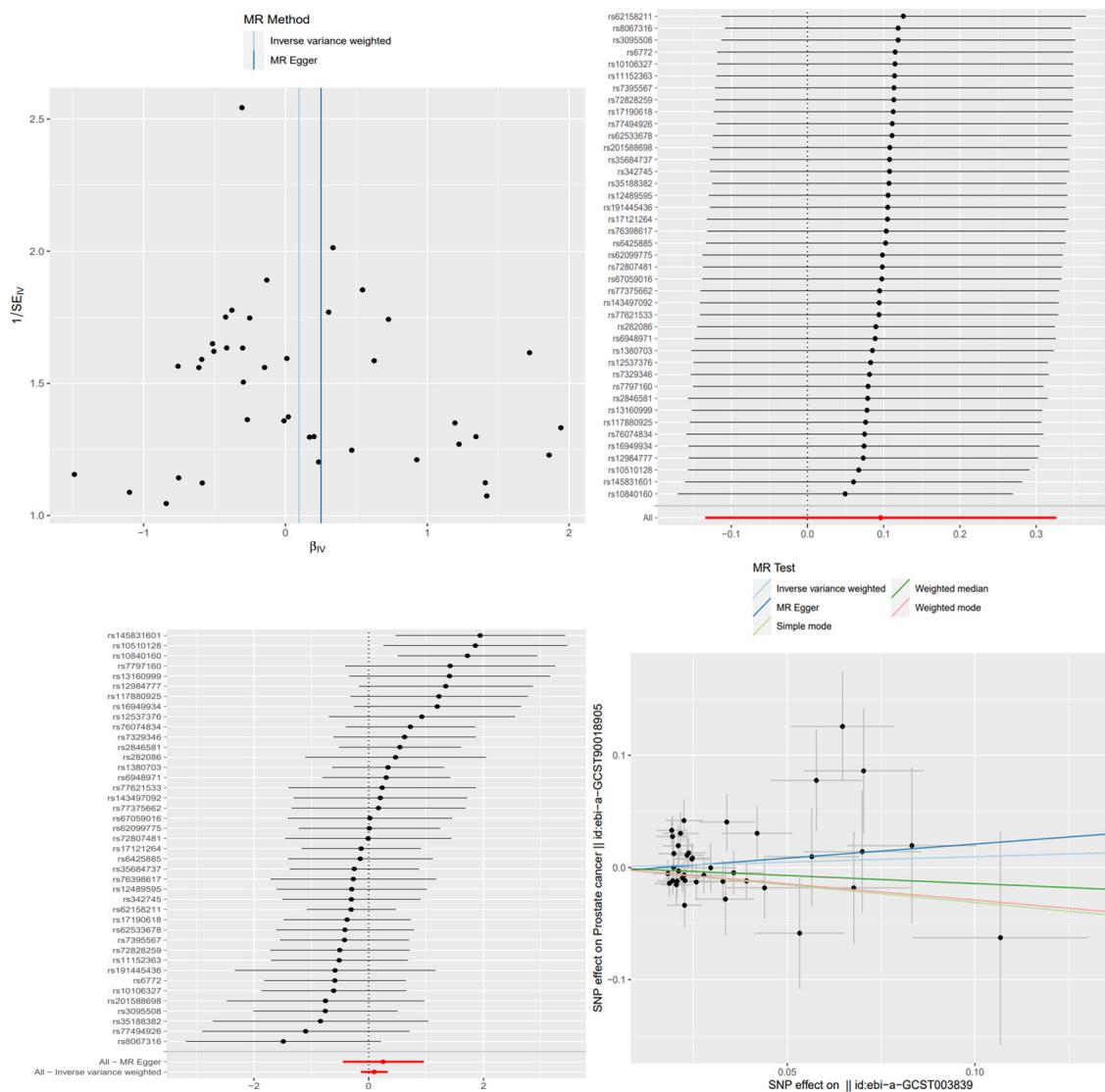


Figure S9 Forest plot, funnel plot, scatter and sensitivity analysis for Sleep duration in prostate cancer (Mendelian Randomization, MR).

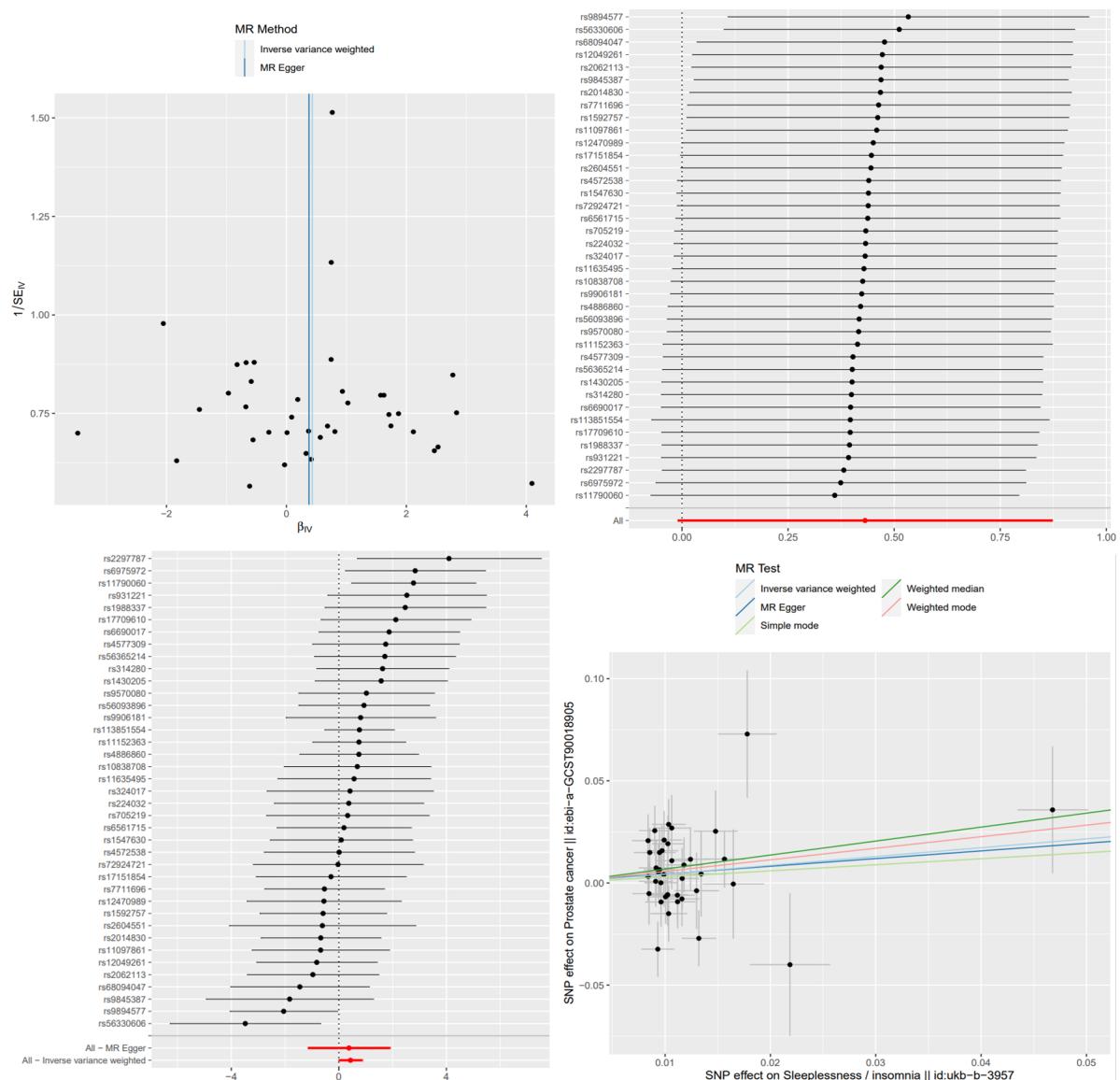


Figure S10 Forest plot, funnel plot, scatter and sensitivity analysis for Sleeplessness/insomnia in prostate cancer (Mendelian Randomization, MR).

Table S1 Baseline characteristics of the study population

| Characteristic | Overall (N=3094) | Prostate Cancer (N=154) | Non-Prostate Cancer (N=2940) | P-Value ¹ |
|----------------------------------------------------|-------------------|-------------------------|------------------------------|----------------------|
| Age (years) | 53 (47, 64) | 72.58 (67, 79) | 53.00 (46, 63) | <0.001 |
| Race | | | | 0.14 |
| Non-Hispanic White | 1,677 (54.2%) | 91 (59.1%) | 1586(53.9%) | |
| Non-Hispanic Black | 640 (20.7%) | 45 (29.2%) | 595(20.2%) | |
| Mexican American | 464 (14.9%) | 8 (5.2%) | 456(15.5%) | |
| Other Hispanic | 313 (10.1%) | 10 (6.5%) | 303(10.3%) | |
| Education_level | | | | 0.5 |
| 9-11th Grade (Includes 12th grade with no diploma) | 461 (14.8%) | 30 (19.5%) | 431 (14.6%) | |
| College Graduate or above | 640 (28%) | 31 (20.1%) | 609 (23.9%) | |
| High School Grad/GED or Equivalent | 736 (23.8%) | 33 (21.4%) | 703 (23.9%) | |
| Less Than 9th Grade | 519 (16.8%) | 21 (13.6%) | 498 (16.9%) | |
| Some College or AA degree | 738 (23.9%) | 39 (25.3%) | 699 (23.8%) | |
| Family_PIR | 3.64 (1.95, 5.00) | 3.12 (2.05, 5.00) | 3.65 (1.95, 5.00) | 0.5 |
| Marital_Status | | | | 0.002 |
| Divorced | 352 (11.3%) | 11 (7.2%) | 341 (11.6%) | |
| Living with partner | 152 (5.0%) | 3 (1.9%) | 149 (5.1%) | |
| Married | 2,075 (67.1%) | 114 (74.0%) | 1,961 (66.7%) | |
| Never married | 214 (6.9%) | 4 (2.6%) | 210 (7.1%) | |
| Separated | 85 (2.7%) | 4 (2.6%) | 81 (2.7%) | |
| Widowed | 216 (7.0%) | 18 (11.7%) | 198 (6.7%) | |
| Annual_Family_Income | | | | 0.013 |
| \$20,001-\$55,000 | 569 (18.3%) | 36 (65%) | 533 (18.8%) | |
| \$55,001-\$75,000 | 139 (4.4%) | 1 (0.5%) | 138 (4.7%) | |
| <\$20,000 | 309 (10.0%) | 7 (9.5%) | 302 (10.2%) | |
| >\$75,001 | 330 (10.7%) | 9 (25%) | 321 (10.9%) | |
| Other | 2316(74.8%) | 101(65.6%) | 2215(75.3%) | |
| Diabetes | | | | 0.5 |
| Diabetes | 514 (16.7%) | 26 (16.9%) | 488 (16.6%) | |
| Non-Diabetes | 2,580 (83.4%) | 128 (83.1%) | 2,452 (83.4%) | |
| Smoke | | | | 0.057 |
| Smoke | 1,936 (62.6%) | 101 (65.6%) | 1,835 (62.4%) | |
| Non-Smoke | 1,158 (37.4%) | 53 (34.4%) | 1,105 (37.6%) | |
| BMI | 28.3 (25.3, 31.6) | 27.7 (25.4, 30.9) | 28.3 (25.3, 31.7) | 0.4 |
| Uric_acid | 357 (315, 410) | 345 (310, 413) | 357 (315, 410) | 0.6 |
| Total_calcium | 2.35 (2.30, 2.43) | 2.35 (2.28, 2.40) | 2.35 (2.30, 2.43) | 0.2 |
| Creatinine | 88 (80, 98) | 97 (81, 107) | 88 (80, 97) | <0.001 |

¹Wilcoxon rank-sum test for complex survey samples; chi-squared test with Rao & Scott's second-order correction; BMI: Body Mass Index.

Table S2 MR result

| X | id.exposure | exposure | id.outcome | outcome | method | nsnp | b | se | Pval | lo_ci | up_ci | or | or_lci95 | or_uci95 |
|---|---------------------|--------------------------------|--------------------|------------------------------------------|---------------------------|------|----------|----------|----------|----------|----------|----------|----------|----------|
| 1 | ebi-a-GCST003839 | Sleep duration | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 41 | 0.251512 | 0.357543 | 0.485956 | -0.44927 | 0.952295 | 1.285968 | 0.638093 | 2.591652 |
| 1 | ebi-a-GCST003839 | Sleep duration | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 41 | -0.14338 | 0.154652 | 0.353866 | -0.4465 | 0.159738 | 0.866424 | 0.639864 | 1.173203 |
| 1 | ebi-a-GCST003839 | Sleep duration | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 41 | 0.095951 | 0.117317 | 0.413427 | -0.13399 | 0.325892 | 1.100705 | 0.874598 | 1.385266 |
| 1 | ebi-a-GCST003839 | Sleep duration | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Simple mode | 41 | -0.31224 | 0.268736 | 0.252171 | -0.83896 | 0.214484 | 0.731807 | 0.432159 | 1.239223 |
| 1 | ebi-a-GCST003839 | Sleep duration | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted mode | 41 | -0.2913 | 0.249065 | 0.24909 | -0.77947 | 0.196864 | 0.747289 | 0.458649 | 1.217579 |
| 2 | ebi-a-GCST006685 | Sleep duration (oversleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 34 | -0.12427 | 0.279104 | 0.659142 | -0.67131 | 0.422775 | 0.883142 | 0.511037 | 1.526191 |
| 2 | ebi-a-GCST006685 | Sleep duration (oversleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 34 | -0.18774 | 0.236071 | 0.426451 | -0.65044 | 0.274957 | 0.828828 | 0.521815 | 1.316474 |
| 2 | ebi-a-GCST006685 | Sleep duration (oversleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 34 | -0.36033 | 0.233338 | 0.122525 | -0.81768 | 0.097007 | 0.697443 | 0.441456 | 1.101868 |
| 2 | ebi-a-GCST006685 | Sleep duration (oversleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Simple mode | 34 | 0.013106 | 0.847347 | 0.987752 | -1.64769 | 1.673907 | 1.013193 | 0.192493 | 5.332961 |
| 2 | ebi-a-GCST006685 | Sleep duration (oversleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted mode | 34 | -0.18823 | 0.204723 | 0.364551 | -0.58948 | 0.213032 | 0.828428 | 0.554614 | 1.237424 |
| 3 | ebi-a-GCST006686 | Sleep duration (undersleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 28 | 0.168604 | 0.603038 | 0.782003 | -1.01335 | 1.350558 | 1.183651 | 0.363001 | 3.859579 |
| 3 | ebi-a-GCST006686 | Sleep duration (undersleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 28 | 0.283672 | 0.415124 | 0.49439 | -0.52997 | 1.097314 | 1.327997 | 0.588622 | 2.996109 |
| 3 | ebi-a-GCST006686 | Sleep duration (undersleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 28 | -0.13755 | 0.294427 | 0.640361 | -0.71463 | 0.439523 | 0.871487 | 0.489372 | 1.551967 |
| 3 | ebi-a-GCST006686 | Sleep duration (undersleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Simple mode | 28 | 0.257898 | 0.804577 | 0.751027 | -1.31907 | 1.834869 | 1.294207 | 0.267383 | 6.264311 |
| 3 | ebi-a-GCST006686 | Sleep duration (undersleepers) | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted mode | 28 | 0.372863 | 0.49305 | 0.456054 | -0.59351 | 1.339242 | 1.451886 | 0.552382 | 3.816148 |
| 4 | finn-b-F5_SLEEP | Sleep disorders | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 16 | -0.01844 | 0.03044 | 0.554379 | -0.0781 | 0.041224 | 0.98173 | 0.92487 | 1.042085 |
| 4 | finn-b-F5_SLEEP | Sleep disorders | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 16 | -0.02119 | 0.027917 | 0.447797 | -0.07591 | 0.033526 | 0.979031 | 0.926901 | 1.034094 |
| 4 | finn-b-F5_SLEEP | Sleep disorders | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 16 | -0.00921 | 0.01851 | 0.618761 | -0.04549 | 0.027068 | 0.990832 | 0.95553 | 1.027438 |
| 4 | finn-b-F5_SLEEP | Sleep disorders | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Simple mode | 16 | -0.02829 | 0.051224 | 0.588827 | -0.12869 | 0.072104 | 0.972102 | 0.879243 | 1.074767 |
| 4 | finn-b-F5_SLEEP | Sleep disorders | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted mode | 16 | -0.03271 | 0.028835 | 0.274465 | -0.08922 | 0.023809 | 0.967821 | 0.91464 | 1.024094 |
| 5 | finn-b-G6_SLEEPAPNO | Sleep apnoea | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 57 | 0.042592 | 0.065414 | 0.517684 | -0.08562 | 0.170804 | 1.043512 | 0.917943 | 1.186259 |
| 5 | finn-b-G6_SLEEPAPNO | Sleep apnoea | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 57 | -0.0034 | 0.037856 | 0.928472 | -0.0776 | 0.0708 | 0.996607 | 0.925338 | 1.073367 |
| 5 | finn-b-G6_SLEEPAPNO | Sleep apnoea | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 57 | 0.02957 | 0.026143 | 0.258013 | -0.02167 | 0.08081 | 1.030012 | 0.978563 | 1.084165 |
| 5 | finn-b-G6_SLEEPAPNO | Sleep apnoea | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Simple mode | 57 | -0.01829 | 0.082761 | 0.825919 | -0.1805 | 0.143924 | 0.981878 | 0.834853 | 1.154796 |
| 5 | finn-b-G6_SLEEPAPNO | Sleep apnoea | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted mode | 57 | -0.02162 | 0.080901 | 0.790297 | -0.18018 | 0.13695 | 0.978615 | 0.835117 | 1.146771 |
| 6 | ukb-b-3957 | Sleeplessness / insomnia | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 205 | -0.10898 | 0.442593 | 0.805751 | -0.97646 | 0.758502 | 0.896748 | 0.376641 | 2.135075 |
| 6 | ukb-b-3957 | Sleeplessness / insomnia | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 205 | 0.368888 | 0.172968 | 0.03295 | 0.02987 | 0.707906 | 1.446125 | 1.03032 | 2.029736 |
| 6 | ukb-b-3957 | Sleeplessness / insomnia | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 205 | 0.297417 | 0.127883 | 0.020035 | 0.046766 | 0.548069 | 1.346377 | 1.047877 | 1.729909 |
| 6 | ukb-b-3957 | Sleeplessness / insomnia | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Simple mode | 205 | 0.78633 | 0.547156 | 0.152215 | -0.2861 | 1.858757 | 2.195326 | 0.751191 | 6.415754 |
| 6 | ukb-b-3957 | Sleeplessness / insomnia | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted mode | 205 | 0.669654 | 0.440633 | 0.130121 | -0.19399 | 1.533295 | 1.953561 | 0.823669 | 4.633418 |
| 7 | ukb-d-20533 | Trouble falling asleep | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 22 | -0.28613 | 0.448598 | 0.530824 | -1.16538 | 0.593126 | 0.751168 | 0.311804 | 1.809637 |
| 7 | ukb-d-20533 | Trouble falling asleep | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 22 | -0.28173 | 0.299568 | 0.346989 | -0.86888 | 0.305425 | 0.754479 | 0.419421 | 1.357202 |
| 7 | ukb-d-20533 | Trouble falling asleep | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 22 | -0.36284 | 0.209511 | 0.083305 | -0.77348 | 0.047806 | 0.6957 | 0.461405 | 1.048967 |
| 7 | ukb-d-20533 | Trouble falling asleep | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Simple mode | 22 | -0.27833 | 0.519935 | 0.598051 | -1.2974 | 0.740739 | 0.757045 | 0.27324 | 2.097485 |
| 7 | ukb-d-20533 | Trouble falling asleep | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted mode | 22 | -0.25899 | 0.491999 | 0.604117 | -1.22331 | 0.705323 | 0.771827 | 0.294254 | 2.0245 |
| 8 | ukb-d-20534 | Sleeping too much | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | MR Egger | 18 | -0.53056 | 0.645905 | 0.423478 | -1.79653 | 0.735414 | 0.588275 | 0.165873 | 2.086345 |
| 8 | ukb-d-20534 | Sleeping too much | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Weighted median | 18 | 0.059111 | 0.329223 | 0.857508 | -0.58617 | 0.704389 | 1.060893 | 0.556456 | 2.02261 |
| 8 | ukb-d-20534 | Sleeping too much | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | Inverse variance weighted | 18 | 0.01533 | 0.288886 | 0 | | | | | |

Table S3 Horizontal pleiotropy in MR

| id.exposure | id.outcome | outcome | exposure | egger_intercept | se | Pval |
|---------------------|--------------------|---------------------------------------------|------------------------|-----------------|----------|----------|
| ebi-a-GCST003839 | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:ebi-a-GCST003839 | -0.00426 | 0.009231 | 0.647258 |
| ebi-a-GCST006685 | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:ebi-a-GCST006685 | -0.00829 | 0.005591 | 0.148017 |
| ebi-a-GCST006686 | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:ebi-a-GCST006686 | -0.00497 | 0.008505 | 0.564229 |
| finn-b-F5_SLEEP | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:finn-b-F5_SLEEP | 0.00366 | 0.009583 | 0.708287 |
| finn-b-G6_SLEEPAPNO | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:finn-b-G6_SLEEPAPNO | -0.00125 | 0.005757 | 0.828883 |
| ukb-b-3957 | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:ukb-b-3957 | 0.003914 | 0.004081 | 0.338625 |
| ukb-d-20533 | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:ukb-d-20533 | -0.00186 | 0.009628 | 0.848606 |
| ukb-d-20534 | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:ukb-d-20534 | 0.015089 | 0.015956 | 0.358378 |
| ukb-e-1220_AFR | ebi-a-GCST90018905 | Prostate cancer id:ebi-a-GCST90018905 | id:ukb-e-1220_AFR | 0.019451 | 0.013652 | 0.169626 |

Table S4 Heterogeneity statistics in MR

| id.exposure | method | Q | Q_df | Q_Pval |
|---------------------|---------------------------|----------|------|----------|
| ebi-a-GCST003839 | MR Egger | 51.38762 | 39 | 0.08843 |
| ebi-a-GCST003839 | Inverse variance weighted | 51.66781 | 40 | 0.102288 |
| ebi-a-GCST006685 | MR Egger | 54.28951 | 32 | 0.008248 |
| ebi-a-GCST006685 | Inverse variance weighted | 58.01768 | 33 | 0.004563 |
| ebi-a-GCST006686 | MR Egger | 36.80182 | 26 | 0.077879 |
| ebi-a-GCST006686 | Inverse variance weighted | 37.28463 | 27 | 0.089876 |
| finn-b-F5_SLEEP | MR Egger | 13.60032 | 14 | 0.479892 |
| finn-b-F5_SLEEP | Inverse variance weighted | 13.74615 | 15 | 0.544863 |
| finn-b-G6_SLEEPAPNO | MR Egger | 52.83696 | 55 | 0.557714 |
| finn-b-G6_SLEEPAPNO | Inverse variance weighted | 52.88412 | 56 | 0.593565 |
| ukb-b-3957 | MR Egger | 270.8554 | 203 | 0.00102 |
| ukb-b-3957 | Inverse variance weighted | 272.0828 | 204 | 0.00101 |
| ukb-d-20533 | MR Egger | 12.894 | 20 | 0.881885 |
| ukb-d-20533 | Inverse variance weighted | 12.9314 | 21 | 0.911008 |
| ukb-d-20534 | MR Egger | 27.13261 | 16 | 0.040033 |
| ukb-d-20534 | Inverse variance weighted | 28.64915 | 17 | 0.037902 |
| ukb-e-1220_AFR | MR Egger | 48.02745 | 20 | 0.000422 |
| ukb-e-1220_AFR | Inverse variance weighted | 52.90255 | 21 | 0.000142 |

Q: Cochran's Q test.