## SUPPLEMENTARY MATERIAL

Table S1. List of the SNPs used for genetic profile risk score for BMI

| SNP | Chr | Position | Genes | BMI- <br> increasing allele | Other allele | иьш и зук 4лп.ь |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| rs1558902 | 16 | 52,361,075 | FTO | A | T | 0.0818 |
| rs6567160 | 18 | 55,980,115 | MC4R | C | T | 0.0556 |
| rs13021737 | 2 | 622,348 | TMEM18 | G | A | 0.0601 |
| rs10938397 | 4 | 44,877,284 | GNPDA2 | G | A | 0.0402 |
| rs543874 | 1 | 176,156,103 | SEC16B | G | A | 0.0482 |
| rs2207139 | 6 | 50,953,449 | TFAP2B | G | A | 0.0447 |
| rs11030104 | 11 | 27,641,093 | BDNF | A | G | 0.0414 |
| rs3101336 | 1 | 72,523,773 | NEGR1 | C | T | 0.0334 |
| rs7138803 | 12 | 48,533,735 | BCDIN3D | A | G | 0.0315 |
| rs10182181 | 2 | 25,003,800 | ADCY3 | G | A | 0.0307 |
| rs3888190 | 16 | 28,796,987 | ATP2A1 | A | C | 0.0309 |
| rs1516725 | 3 | 187,306,698 | ETV5 | C | T | 0.0451 |
| rs12446632 | 16 | 19,842,890 | GPRC5B | G | A | 0.0403 |
| rs2287019 | 19 | 50,894,012 | QPCTL | C | T | 0.0360 |
| rs16951275 | 15 | 65,864,222 | MAP2K5 | T | C | 0.0311 |
| rs3817334 | 11 | 47,607,569 | MTCH2 | T | C | 0.0262 |
| rs2112347 | 5 | 75,050,998 | POC5 | T | G | 0.0261 |
| rs12566985 | 1 | 74,774,781 | FPGT | G | A | 0.0242 |
| rs3810291 | 19 | 52,260,843 | ZC3H4 | A | G | 0.0283 |
| rs7141420 | 14 | 78,969,207 | NRXN3 | T | C | 0.0235 |
| rs13078960 | 3 | 85,890,280 | CADM2 | G | T | 0.0297 |
| rs10968576 | 9 | 28,404,339 | LINGO2 | G | A | 0.0249 |
| rs17024393 | 1 | 109,956,211 | GNAT2 | C | T | 0.0658 |
| rs12429545 | 13 | 53,000,207 | OLFM4 | A | G | 0.0334 |
| rs13107325 | 4 | 103,407,732 | SLC39A8 | T | C | 0.0477 |
| rs11165643 | 1 | 96,696,685 | PTBP2 | T | C | 0.0218 |
| rs17405819 | 8 | 76,969,139 | HNF4G | T | C | 0.0224 |
| rs1016287 | 2 | 59,159,129 | LINC01122 | T | C | 0.0229 |
| rs4256980 | 11 | 8,630,515 | TRIM66 | G | C | 0.0209 |
| rs12401738 | 1 | 78,219,349 | FUBP1 | A | G | 0.0211 |
| rs205262 | 6 | 34,671,142 | C6orf106 | G | A | 0.0221 |
| rs12016871 | 13 | 26,915,782 | MTIF3 | T | C | 0.0298 |
| rs12940622 | 17 | 76,230,166 | RPTOR | G | A | 0.0182 |
| rs11847697 | 14 | 29,584,863 | PRKD1 | T | C | 0.0492 |
| rs2075650 | 19 | 50,087,459 | TOMM40 | A | G | 0.0258 |
| rs2121279 | 2 | 142,759,755 | LRP1B | T | C | 0.0245 |
| rs29941 | 19 | 39,001,372 | KCTD15 | G | A | 0.0182 |
| rs6091540 | 20 | 50,521,269 | ZFP64 | C | T | 0.0188 |
| rs7715256 | 5 | 153,518,086 | GALNT10 | G | T | 0.0163 |
| rs2176040 | 2 | 226,801,046 | LOC646736 | A | G | 0.0141 |
| rs657452 | 1 | 49,362,434 | AGBL4 | A | G | 0.0227 |
| rs12286929 | 11 | 114,527,614 | CADM1 | G | A | 0.0217 |
| rs7903146 | 10 | 114,748,339 | TCF7L2 | C | T | 0.0234 |
| rs10132280 | 14 | 24,998,019 | STXBP6 | C | A | 0.0230 |
| rs17094222 | 10 | 102,385,430 | HIF1AN | C | T | 0.0249 |
| rs7599312 | 2 | 213,121,476 | ERBB4 | G | A | 0.0220 |
| rs2365389 | 3 | 61,211,502 | FHIT | C | T | 0.0200 |


| rs2820292 | 1 | 200,050,910 | NAV1 | C | A | 0.0195 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| rs12885454 | 14 | 28,806,589 | PRKD1 | C | A | 0.0207 |
| rs16851483 | 3 | 142,758,126 | RASA2 | T | G | 0.0483 |
| rs1167827 | 7 | 75,001,105 | HIP1 | G | A | 0.0202 |
| rs758747 | 16 | 3,567,359 | NLRC3 | T | C | 0.0225 |
| rs1928295 | 9 | 119,418,304 | TLR4 | T | C | 0.0188 |
| rs9925964 ${ }^{1}$ | 16 | 31,037,396 | KAT8 | A | G | 0.0192 |
| rs11126666 | 2 | 26,782,315 | KCNK3 | A | G | 0.0207 |
| rs2650492 | 16 | 28,240,912 | SBK1 | A | G | 0.0207 |
| rs6804842 | 3 | 25,081,441 | RARB | G | A | 0.0185 |
| rs4740619 | 9 | 15,624,326 | C9orf93 | T | C | 0.0179 |
| rs13191362 | 6 | 162,953,340 | PARK2 | A | G | 0.0277 |
| rs3736485 | 15 | 49,535,902 | DMXL2 | A | G | 0.0176 |
| rs17001654 ${ }^{2}$ | 4 | 77,348,592 | SCARB2 | G | C | 0.0306 |
| rs11191560 | 10 | 104,859,028 | NT5C2 | C | T | 0.0308 |
| rs1528435 | 2 | 181,259,207 | UBE2E3 | T | C | 0.0178 |
| rs1000940 | 17 | 5,223,976 | RABEP1 | G | A | 0.0192 |
| rs2033529 ${ }^{1}$ | 6 | 40,456,631 | TDRG1 | G | A | 0.0190 |
| rs11583200 | 1 | 50,332,407 | ELAVL4 | C | T | 0.0177 |
| rs9400239 | 6 | 109,084,356 | FOXO3 | C | T | 0.0188 |
| rs10733682 | 9 | 128,500,735 | LMX1B | A | G | 0.0174 |
| rs11688816 | 2 | 62,906,552 | EHBP1 | G | A | 0.0172 |
| rs11057405 | 12 | 121,347,850 | CLIP1 | G | A | 0.0307 |
| rs11727676 | 4 | 145,878,514 | HHIP | T | C | 0.0358 |
| rs3849570 | 3 | 81,874,802 | GBE1 | A | C | 0.0188 |
| rs6477694 | 9 | 110,972,163 | EPB41L4B | C | T | 0.0174 |
| rs7899106 | 10 | 87,400,884 | GRID1 | G | A | 0.0395 |
| rs2176598 | 11 | 43,820,854 | HSD17B12 | T | C | 0.0198 |
| rs2245368 | 7 | 76,446,079 | DTX2P1 | C | T | 0.0317 |
| rs17724992 | 19 | 18,315,825 | PGPEP1 | A | G | 0.0194 |
| rs7243357 | 18 | 55,034,299 | GRP | T | G | 0.0217 |
| rs1808579 | 18 | 19,358,886 | C18orf8 | C | T | 0.0167 |
| rs2033732 | 8 | 85,242,264 | RALYL | C | T | 0.0192 |
| rs1441264 | 13 | 78,478,920 | MIR548A2 | A | G | 0.0175 |
| rs2080454 | 16 | 47,620,091 | CBLN1 | C | A | 0.0168 |
| rs7164727 | 15 | 70,881,044 | LOC100287559 | T | C | 0.0180 |
| rs17203016 | 2 | 207,963,763 | CREB1 | G | A | 0.0210 |
| rs977747 | 1 | 47,457,264 | TAL1 | T | G | 0.0167 |
| rs9914578 | 17 | 1,951,886 | SMG6 | G | C | 0.0201 |
| rs9374842 | 6 | 120,227,364 | LOC285762 | T | C | 0.0187 |
| rs16907751 | 8 | 81,538,012 | ZBTB10 | C | T | 0.0350 |
| rs9540493 | 13 | 65,103,705 | MIR548X2 | A | G | 0.0172 |
| rs7239883 | 18 | 38,401,669 | LOC284260 | G | A | 0.0164 |
| rs13201877 | 6 | 137,717,234 | IFNGR1 | G | A | 0.0233 |
| rs2836754 | 21 | 39,213,610 | ETS2 | C | T | 0.0164 |
| rs492400 | 2 | 219,057,996 | USP37 | C | T | 0.0158 |
| rs9641123 | 7 | 93,035,668 | CALCR | C | G | 0.0191 |
| rs1460676 | 2 | 164,275,935 | FIGN | C | T | 0.0197 |
| rs4787491 | 16 | 29,922,838 | INO80E | G | A | 0.0159 |
| rs6465468 | 7 | 95,007,450 | ASB4 | T | G | 0.0166 |

[^0]Table S2. Cohort characteristic by genetic risk score quartiles by combined categories of total self-reported PA and total sedentary behaviour

|  | $\begin{gathered} \text { Active / } \\ \text { Low sedentary } \\ \text { behaviour } \end{gathered}$ | Active / <br> High sedentary behaviour | Inactive / Low sedentary behaviour | Inactive / <br> High sedentary behaviour |
| :---: | :---: | :---: | :---: | :---: |
| Socio-demographics |  |  |  |  |
| Total n | 111,565 | 72,403 | 83,714 | 70,354 |
| Women, n (\%) | 58,138 (52.1) | 37,029 (51.1) | 47,678 (57.0) | 38,809 (55.0) |
| Age (years) | 56.9 (8.3) | 56.6 (7.9) | 57.1 (7.9) | 56.8 (7.6) |
| Deprivation index tertile Lower <br> Middle <br> Higher | $\begin{aligned} & 41,445(37.2) \\ & 38,707(34.7) \\ & 31,293(28.1) \\ & \hline \end{aligned}$ | $\begin{aligned} & 26,711(36.9) \\ & 25,243(34.9) \\ & 20,361(28.2) \\ & \hline \end{aligned}$ | $29,661(35.5)$ $28,094(33.6)$ $25,867(30.9)$ | $\begin{aligned} & 23,686(33.6) \\ & 23,922(34.0) \\ & 22,819(32.4) \\ & \hline \end{aligned}$ |
| Smoking status, n (\%) <br> Never <br> Previous <br> Current | $\begin{gathered} 63,893 \text { (57.4) } \\ 38,171 \text { (34.3) } \\ 9,214(8.3) \\ \hline \end{gathered}$ | $\begin{gathered} 37,303(51,7) \\ 27,618(38.2) \\ 7,284(10.1) \\ \hline \end{gathered}$ | $\begin{gathered} 47,932(57.6) \\ 27,026(32.4) \\ 8,316(10.0) \\ \hline \end{gathered}$ | $\begin{gathered} 35,182(50.1) \\ 25,991(37.0) \\ 9,097(12.9) \\ \hline \end{gathered}$ |
| Obesity-related markers |  |  |  |  |
| BMI kg.m ${ }^{-2}$ | 26.3 (4.1) | 27.8 (4.5) | 27.2 (4.8) | 29.0 (5.4) |
| BMI Categories, n (\%) <br> Underweight (<18.5) <br> Normal weight (18.5-24.9) <br> Overweight (25.0 to 29.9) <br> Obese ( $\geq 30.0$ ) | $\begin{gathered} 671(0.6) \\ 45,924(41.2) \\ 47,294(42.4) \\ 17,676(15.8) \end{gathered}$ | $\begin{gathered} 224(0.3) \\ 20,332(28.1) \\ 32,972(45.5) \\ 18,875(26.1) \end{gathered}$ | $\begin{gathered} 532(0.6) \\ 28,584(34.1) \\ 35,106(42.0) \\ 19,492(23.3) \end{gathered}$ | $\begin{gathered} 273(0.4) \\ 15,860(22.5) \\ 29,039(41.2) \\ 25,362(35.9) \end{gathered}$ |
| Body fat (\%) | 29.3 (8.3) | 31.5 (8.2) | 31.8 (8.5) | 33.9 (8.4) |
| Waist Circumference (cm) | 87.4 (12.3) | 91.1 (13.2) | 90.1 (13.4) | 94.4 (14.4) |
| Central Obesity, n (\%) | 26,150 (23.4) | 24,941 (34.4) | 28,667 (34.2) | 33,205 (47.1) |
| Physical activity |  |  |  |  |
| Total PA (METs-hr.week ${ }^{-1}$ ), mean (SD) | 72.3 (73.6) | 69.6 (72.0) | 13.6 (18.8) | 12.0 (17.3) |
| Objective total PA (milli-gravity.day ${ }^{1}$ ), mean (SD) | 30.1 (8.8) | 27.9 (8.2) | 27.1 (7.4) | 24.9 (7.0) |
| TV viewing ( ${\mathrm{h} . \mathrm{day}^{-1} \text { ) }}^{\text {d }}$ | 2.0 (1.1) | 3.7 (1.5) | 2.1 (1.1) | 4.0 (1.7) |
| Total Sedentary Behaviour (h.day ${ }^{-1}$ ) | 3.6 (1.1) | 6.9 (1.8) | 3.6 (1.1) | 7.1 (2.0) |
| Dietary intake |  |  |  |  |
| Total energy intake ( $\mathrm{Kcal}^{\text {d }}$ day ${ }^{-1}$ ) | 2,212 (656) | 2,180 (668) | 2,149 (620) | 2,131 (647) |
| Protein intake (\% of TE) | 15.4 (3.4) | 15.6 (3.6) | 15.4 (3.4) | 15.6 (3.7) |
| Carbohydrates intake (\% of TE) | 47.5 (7.9) | 47.0 (8.0) | 47.1 (7.9) | 46.6 (8.1) |
| Total Fat intake (\% of TE) | 31.7 (6.6) | 32.0 (6.7) | 32.3 (6.5) | 32.6 (6.8) |
| Saturated intake (\% of TE) | 12.1 (3.3) | 12.3 (3.3) | 12.4 (3.3) | 12.6 (3.4) |
| Polyunsaturated fat intake (\% of TE) | 14.5 (7.2) | 14.5 (7.4) | 14.2 (7.1) | 14.4 (7.4) |
| Processed meat intake, $\mathrm{n}(\%)$ Never <br> <1 a week <br> 2-4 a week <br> $>5$ times a week | $\begin{gathered} 11,338(10.2) \\ 34,799(31.2) \\ 61,623(55.3) \\ 3,724(3.3) \end{gathered}$ | $\begin{gathered} 5,377(7.4) \\ 20,981(29.0) \\ 43,015(59.4) \\ 3,000(4.2) \end{gathered}$ | $\begin{gathered} 6,795(8.1) \\ 25,394(30.4) \\ 48,067(57.6) \\ 3,226(3.9) \end{gathered}$ | $\begin{gathered} 4,241(6.0) \\ 20,034(28.5) \\ 42,864(60.9) \\ 3,258(4.6) \end{gathered}$ |
| Sugar intake (\% of TE) | 22.9 (6.8) | 22.6 (6.9) | 22.2 (6.7) | 21.9 (7.0) |
| Starch intake (g.day ${ }^{-1}$ ) | 124.5 (46.0) | 122.0 (47.2) | 122.6 (44.5) | 121.2 (46.6) |
| Alcohol intake (\% of TE) | 5.4 (6.4) | 5.4 (6.6) | 5.2 (6.6) | 5.2 (6.8) |
| Health status, n (\%) |  |  |  |  |
| Diabetes history | 3,547 (3.2) | 3,534 (4.9) | 3,878 (4.6) | 5,185 (7.4) |
| Cancer history | 8,285 (7.4) | 5,569 (7.7) | 6,642 (8.0) | 5,971 (8.5) |
| CVDs | 28,754 (25.8) | 21,830 (30.2) | 25,441 (30.4) | 24,854 (35.2) |
| Hypertension | 22,580 (20.2) | 16,752 (23.1) | 19,540 (23.3) | 18,687 (26.5) |

Data presented as mean and SD for continuous variables and as $n$ and \% for categorical variables.
TE: \% of total energy intake; CVD: cardiovascular diseases, MET: metabolic-equivalent task; BMI: body mass index.

Table S3. Cohort characteristic by genetic risk score quartiles by combined categories of total objective PA and total sedentary behaviour

|  | Active / Low sedentary behaviour | Active / High sedentary behaviour | Inactive / Low sedentary behaviour | Inactive / High sedentary behaviour |
| :---: | :---: | :---: | :---: | :---: |
| Socio-demographics |  |  |  |  |
| Total n | 22,045 | 11,737 | 18,265 | 15,991 |
| Women, n (\%) | 12,758 (57.9) | 6,964 (59.3) | 9,515 (52.1) | 8,427 (52.7) |
| Age (years) | 54.8 (7.8) | 54.5 (7.4) | 58.3 (7.5) | 58.0 (7.2) |
| Deprivation index tertile Lower <br> Middle <br> Higher | $\begin{aligned} & 8,856(40.2) \\ & 7,604(34.5) \\ & 5,568(25.3) \end{aligned}$ | 4,567 (39.0) 4,203 (35.8) 2,955 (25.2) | $\begin{aligned} & 7,182(39.4) \\ & 6,300(34.6) \\ & 4,755(26.0) \\ & \hline \end{aligned}$ | $\begin{aligned} & 6,099(38.2) \\ & 5,687(35.6) \\ & 4,187(26.2) \\ & \hline \end{aligned}$ |
| Smoking status, n (\%) <br> Never <br> Previous <br> Current | $\begin{gathered} 13,519(61.5) \\ 7,316(33.3) \\ 1,161(5.2) \\ \hline \end{gathered}$ | $\begin{gathered} 6,506(55.5) \\ 4,425(37.8) \\ 785(6.7) \\ \hline \end{gathered}$ | $\begin{gathered} 10,722(58.8) \\ 6,338(34.8) \\ 1,160(6.4) \\ \hline \end{gathered}$ | $\begin{gathered} 8,232(51.6) \\ 6,330(39.7) \\ 1,396(8.7) \\ \hline \end{gathered}$ |
| Obesity-related markers |  |  |  |  |
| BMI kg.m ${ }^{-2}$ | 25.2 (3.7) | 26.6 (4.2) | 26.9 (4.4) | 28.6 (5.1) |
| BMI Categories, n (\%) <br> Underweight (<18.5) <br> Normal weight (18.5-24.9) <br> Overweight (25.0 to 29.9) <br> Obese ( $\geq 30.0$ ) | $\begin{gathered} 186(0.8) \\ 11,483(52.1) \\ 8,238(37.4) \\ 2,138(9.7) \\ \hline \end{gathered}$ | $\begin{gathered} 60(0.5) \\ 4,520(38.5) \\ 5,069(43.2) \\ 2,088(17.8) \\ \hline \end{gathered}$ | $\begin{gathered} 89(0.5) \\ 6,569(36.0) \\ 7,952(43.5) \\ 3,655(20.0) \\ \hline \end{gathered}$ | $\begin{gathered} 36(0.2) \\ 3,884(24.3) \\ 6,872(43.0) \\ 5,199(32.5) \\ \hline \end{gathered}$ |
| Body fat (\%) | 28.3 (8.0) | 30.7 (8.0) | 30.9 (8.4) | 33.2 (8.4) |
| Waist Circumference (cm) | 84.2 (11.5) | 87.2 (12.4) | 89.8 (12.7) | 93.6 (13.8) |
| Central Obesity, n (\%) | 3,563 (16.2) | 3,023 (25.8) | 5,593 (30.6) | 6,921 (43.3) |
| Physical activity |  |  |  |  |
| Total PA (METs-hr.week ${ }^{-1}$ ), mean (SD) | 50.0 (58.0) | 48.4 (59.8) | 37.2 (48.4) | 33.4 (46.2) |
| Objective total PA (milli-gravity.day ${ }^{-}$ <br> ${ }^{1}$ ), mean (SD) | 34.5 (7.0) | 33.6 (6.1) | 22.2 (3.5) | 21.4 (3.9) |
| Physical active individuals, n (\%) | 14,596 (66.2) | 7,315 (62.3) | 9,651 (52.8) | 7,510 (47.0) |
| TV viewing (h.day ${ }^{-1}$ ) | 1.7 (1.0) | 3.3 (1.4) | 1.9 (1.0) | 3.7 (1.5) |
| Total Sedentary Behaviour (h.day ${ }^{-1}$ ) | 3.5 (1.0) | 6.7 (1.8) | 3.7 (1.0) | 7.0 (1.9) |
| Dietary intake |  |  |  |  |
| Total energy intake ( $\mathrm{Kcal}^{\text {d }} \mathrm{day}^{-1}$ ) | 2,247 (628) | 2,215 (636) | 2,140 (582) | 2,136 (597) |
| Protein intake (\% of TE) | 15.1 (3.0) | 15.4 (3.3) | 15.5 (3.2) | 15.6 (3.4) |
| Carbohydrates intake (\% of TE) | 47.5 (7.4) | 47.0 (7.7) | 47.0 (7.6) | 46.6 (7.6) |
| Total Fat intake (\% of TE) | 32.2 (6.1) | 32.3 (6.3) | 32.2 (6.3) | 32.4 (6.3) |
| Saturated intake (\% of TE) | 12.3 (3.1) | 12.4 (3.1) | 12.4 (3.2) | 12.6 (3.2) |
| Polyunsaturated fat intake (\% of TE) | 14.9 (6.8) | 14.9 (7.0) | 14.1 (6.5) | 14.2 (6.7) |
| Processed meat intake, $\mathrm{n}(\%)$ <br> Never <br> <1 a week <br> 2-4 a week <br> >5 times a week | $\begin{gathered} 2,640(12.0) \\ 7,206(32.7) \\ 11,506(52.2) \\ 680(3.1) \end{gathered}$ | $\begin{gathered} 1,079(9.2) \\ 3,686(31.4) \\ 6,556(55.9) \\ 410(3.5) \\ \hline \end{gathered}$ | $\begin{gathered} 1,635(8.9) \\ 5,916(32.4) \\ 9,997(54.8) \\ 707(3.9) \\ \hline \end{gathered}$ | $\begin{gathered} 1,074(6.7) \\ 4,927(30.8) \\ 9,297(58.2) \\ 685(4.3) \\ \hline \end{gathered}$ |
| Sugar intake (\% of TE) | 22.8 (6.3) | 22.6 (6.5) | 22.3 (6.4) | 22.0 (6.6) |
| Starch intake (g.day ${ }^{-1}$ ) | 127.1 (43.9) | 124.2 (44.6) | 120.9 (41.2) | 120.3 (42.7) |
| Alcohol intake (\% of TE) | 5.2 (5.8) | 5.2 (6.1) | 5.3 (6.2) | 5.3 (6.4) |
| Health status, n (\%) |  |  |  |  |
| Diabetes history | 293 (1.3) | 283 (2.4) | 683 (3.7) | 1,009 (6.3) |
| Cancer history | 1,358 (6.2) | 774 (6.6) | 1,528 (8.4) | 1,365 (8.6) |
| CVDs | 3,861 (17.5) | 2,395 (20.4) | 5,430 (29.7) | 5,507 (34.4) |
| Hypertension | 3,180 (14.4) | 1,946 (16.6) | 4,332 (23.7) | 4,328 (27.1) |

Data presented as mean and SD for continuous variables and as n and \% for categorical variables.
TE: \% of total energy intake; CVD: cardiovascular diseases, MET: metabolic-equivalent task; BMI: body mass index.

Table S4. Cohort characteristic by genetic risk score quartiles by combined categories of total self-reported PA and total TV-viewing

|  | Active / <br> Low sedentary <br> behaviour | Active / <br> High sedentary <br> behaviour | Inactive / <br> Low sedentary <br> behaviour | Inactive / <br> High sedentary <br> behaviour |
| :--- | :---: | :---: | :---: | :---: |
| Socio-demographics |  |  |  |  |
| Total n | 112,938 | 58,611 | 98,930 | 65,771 |
| Women, n (\%) | $61,133(54.1)$ | $30,605(52.28)$ | $52,536(53.1)$ | $35,936(54.6)$ |
| Age (years) | $56.9(7.9)$ | $56.6(8.4)$ | $57.0(7.7)$ | $56.8(8.2)$ |
| Deprivation index tertile |  |  |  |  |
| Lower | $42,742(37.9)$ | $19,236(32.8)$ | $38,298(38.8)$ | $20,731(31.5)$ |
| Middle | $39,537(35.0)$ | $20,005(34.2)$ | $34,295(34.7)$ | $21,583(32.9)$ |
| Higher | $30,525(27.1)$ | $19,308(33.0)$ | $26,212(26.5)$ | $23,371(35.6)$ |
| Smoking status, n (\%) |  |  |  |  |
| Never | $64,033(56.8)$ | $29,980(51.3)$ | $56,686(57.5)$ | $32,586(49.8)$ |
| Previous | $39,620(35.2)$ | $21,457(36.7)$ | $33,503(34.0)$ | $23,623(36.1)$ |
| Current | $9,003(8.0)$ | $7,009(12.0)$ | $8,397(8.5)$ | $9,244(14.1)$ |
| Obesity-related markers |  |  |  |  |
| BMI kg.m |  |  |  | $27.3(4.7)$ |

Data presented as mean and SD for continuous variables and as $n$ and \% for categorical variables.
TE: \% of total energy intake; CVD: cardiovascular diseases, MET: metabolic-equivalent task; BMI: body mass index.

Table S5. Cohort characteristic by genetic risk score quartiles by combined categories of total objective PA and total TV-viewing

|  | Active / Low sedentary behaviour | Active / <br> High sedentary behaviour | Inactive / Low sedentary behaviour | Inactive / High sedentary behaviour |
| :---: | :---: | :---: | :---: | :---: |
| Socio-demographics |  |  |  |  |
| Total n | 25,343 | 8,393 | 22,602 | 11,575 |
| Women, n (\%) | 14,905 (58.8) | 4,778 (56.9) | 11,729 (51.9) | 6,146 (53.1) |
| Age (years) | 55.0 (7.6) | 53.8 (7.8) | 58.4 (7.2) | 57.6 (7.8) |
| Deprivation index tertile Lower <br> Middle <br> Higher | $\begin{gathered} 10,284(40.6) \\ 8,801(34.8) \\ 6,235(24.6) \\ \hline \end{gathered}$ | $\begin{aligned} & 3,114 \text { (37.1) } \\ & 2,999(35.8) \\ & 2,274(27.1) \end{aligned}$ | $\begin{aligned} & 8,936(39.6) \\ & 7,965(35.3) \\ & 5,670(25.1) \\ & \hline \end{aligned}$ | $\begin{aligned} & 4,318(37.4) \\ & 3,998(34.5) \\ & 3,244(28.1) \end{aligned}$ |
| Smoking status, n (\%) Never Previous Current | $\begin{gathered} 15,300(60.5) \\ 8,698(34.4) \\ 1,294(5.1) \\ \hline \end{gathered}$ | $\begin{gathered} 4,696(56.1) \\ 3,029(36.2) \\ 650(7.7) \\ \hline \end{gathered}$ | $\begin{gathered} 12,933(57.3) \\ 8,154(36.2) \\ 1,466(6.5) \\ \hline \end{gathered}$ | $\begin{gathered} 5,970(51.7) \\ 4,491(38.9) \\ 1,089(9.4) \\ \hline \end{gathered}$ |
| Obesity-related markers |  |  |  |  |
| BMI kg.m ${ }^{-2}$ | 25.3 (3.7) | 26.7 (4.2) | 27.2 (4.6) | 28.7 (5.1) |
| BMI Categories, n (\%) <br> Underweight (<18.5) <br> Normal weight (18.5-24.9) <br> Overweight (25.0 to 29.9) <br> Obese ( $\geq 30.0$ ) | $\begin{gathered} 205(0.8) \\ 12,928(51.1) \\ 9,542(37.6) \\ 2,668(10.5) \end{gathered}$ | $\begin{gathered} 41(0.4) \\ 3,060(36.5) \\ 3,741(44.6) \\ 1,551(18.5) \end{gathered}$ | $\begin{gathered} 97(0.4) \\ 7,702(34.1) \\ 9,828(43.5) \\ 4,975(22.0) \end{gathered}$ | $\begin{gathered} 28(0.2) \\ 2,727(23.6) \\ 4,970(42.9) \\ 3,850(33.3) \end{gathered}$ |
| Body fat (\%) | 28.6 (8.0) | 30.6 (8.1) | 31.2 (8.4) | 33.5 (8.5) |
| Waist Circumference (cm) | 84.4 (11.6) | 87.7 (12.4) | 90.5 (13.0) | 93.7 (13.8) |
| Central Obesity, n (\%) | 4,364 (17.2) | 2,206 (26.3) | 7,409 (32.8) | 5,067 (43.8) |
| Physical activity |  |  |  |  |
| Total PA (METs-hr.week ${ }^{-1}$ ), mean (SD) | 49.4 (57.4) | 49.6 (62.2) | 36.0 (47.6) | 34.4 (47.1) |
| Objective total PA (milli-gravity.day ${ }^{1}$ ), mean (SD) | 34.4 (6.8) | 33.7 (6.3) | 22.0 (3.6) | 21.3 (3.9) |
| Physical active individuals, n (\%) | 16,761 (66.1) | 5,134 (61.2) | 11,700 (51.2) | 5,437 (47.0) |
| TV viewing ( $\mathrm{h}^{\text {day }}{ }^{-1}$ ) | 1.6 (0.9) | 4.0 (1.0) | 1.9 (0.9) | 4.4 (1.2) |
|  | 4.0 (1.7) | 6.4 (1.9) | 4.5 (1.8) | 6.9 (2.0) |
| Dietary intake |  |  |  |  |
| Total energy intake ( $\mathrm{Kcal}^{\text {d }}$ day ${ }^{-1}$ ) | 2,237 (624) | 2,232 (654) | 2,136 (585) | 2,142 (600) |
| Protein intake (\% of TE) | 15.2 (3.1) | 15.4 (3.4) | 15.5 (3.2) | 15.7 (3.5) |
| Carbohydrates intake (\% of TE) | 47.5 (7.4) | 47.0 (7.6) | 46.8 (7.6) | 46.7 (7.6) |
| Total Fat intake (\% of TE) | 32.1 (6.2) | 32.5 (6.3) | 32.2 (6.3) | 32.5 (6.4) |
| Saturated intake (\% of TE) | 12.3 (3.1) | 12.5 (3.1) | 12.4 (3.2) | 12.6 (3.2) |
| Polyunsaturated fat intake (\% of TE) | 14.8 (6.8) | 15.2 (7.2) | 14.1 (6.5) | 14.4 (6.8) |
| Processed meat intake, $\mathrm{n}(\%)$ <br> Never <br> <1 a week <br> 2-4 a week <br> $>5$ times a week | $\begin{gathered} 3,031(12.0) \\ 8,376(33.1) \\ 13,145(51.9) \\ 783(3.0) \end{gathered}$ | $\begin{gathered} 687(8.2) \\ 2,496(29.8) \\ 4,899(58.4) \\ 303(3.6) \end{gathered}$ | $\begin{gathered} 1,993(8.8) \\ 7,437(32.9) \\ 12,317(54.5) \\ 848(3.8) \\ \hline \end{gathered}$ | $\begin{gathered} 712 \text { (6.2) } \\ 3,385(29.3) \\ 6,932(59.9) \\ 537(4.6) \end{gathered}$ |
| Sugar intake (\% of TE) | 22.8 (6.3) | 22.2 (6.6) | 22.3 (6.4) | 21.9 (6.6) |
| Starch intake (g.day ${ }^{-1}$ ) | 126.1 (43.9) | 126.2 (45.1) | 120.1 (41.1) | 121.8 (43.4) |
| Alcohol intake (\% of TE) | 5.2 (5.8) | 5.2 (6.2) | 5.4 (6.2) | 5.1 (6.6) |
| Health status, n (\%) |  |  |  |  |
| Diabetes history | 373 (1.5) | 201 (2.4) | 924 (4.1) | 765 (6.6) |
| Cancer history | 1,616 (6.4) | 514 (6.1) | 1,896 (8.4) | 986 (8.5) |
| CVDs | 4,497 (17.7) | 1,749 (20.8) | 6,785 (30.0) | 4,112 (35.5) |
| Hypertension | 3,687 (14.6) | 1,430 (17.0) | 5,434 (24.0) | 3,195 (27.6) |

Data presented as mean and SD for continuous variables and as n and \% for categorical variables.
TE: \% of total energy intake; CVD: cardiovascular diseases, MET: metabolic-equivalent task; BMI: body mass index.

Table S6. Correlation matrix between physical activity, sedentary-related behaviours and genetic profile risk score for BMI.

| Exposures | Self-reported <br> PA | Objective PA | Sedentary <br> behaviours | TV-viewing |
| :--- | :---: | :---: | :--- | :--- |
| Self-reported PA |  |  |  |  |
| Objective PA | $0.180^{*}$ |  |  |  |
| Sedentary behaviours | -0.044 | $-0.183^{*}$ |  |  |
| TV-viewing | -0.036 | $-0.198^{*}$ | $0.644^{* *}$ |  |
| GPRS-BMI | 0.002 | -0.009 | 0.017 | 0.016 |

Pearson correlation coefficient. P-values at *<0.01, ${ }^{* *}<0.0001$

Table S7. Association of genetic profile risk score with body mass index, overweight and obesity.

|  |  | BMI (kg.m $\left.{ }^{-2}\right)^{*}$ |  | BMI $\mathbf{2} \mathbf{2 5}^{\text { }}$ |  | BMI $\geq 30^{\ddagger}$ |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Models | N | Beta (95\% CI) | $p$-value | OR (95\% CI) | $p$-value | OR (95\% CI) | $p$-value |
| Model 1 | 338,216 | 0.54 (0.53; 0.56) | $2.1 \times 10^{-241}$ | 1.18 (1.16; 1.19) | $1.9 \times 10^{-281}$ | 1.30 (1.28; 1.31) | $1.7 \times 10^{-253}$ |
| Model 2 | 229,871 | 0.51 (0.49; 0.53) | $4.3 \times 10^{-207}$ | 1.17 (1.16; 1.18) | $2.0 \times 10^{-193}$ | 1.29 (1.28; 1.31) | $1.5 \times 10^{-191}$ |

Data presented as beta coefficients $\left(^{*}\right)$ or odds ratio (OR) $\left(^{\ddagger}\right.$ ) and the corresponding $95 \% \mathrm{CI}$. *The beta coefficient indicates the change in BMI per SD increase in GPRS. ${ }^{\ddagger}$ The OR indicates the odds ratio for extra risk of being overweight (BMI $\geq 25.0$ ) or obese ( $\mathrm{BMI} \geq 30.0$ ) per SD increase in GPRS.
Analyses for model 1 were adjusted for age, sex, deprivation, education qualifications, recruitment center, month of recruitment, the first 10 principal components of ancestry and genotyping batch, smoking status, dietary intake (alcohol, fruit \& vegetable, red meat, processed meat, cereals, bread and cheese) and comorbidities (diabetes, hypertension, cardiovascular diseases and cancer). Analyses performed for objectively measured PA were additionally adjusted for season and wearing time. Analyses for model 2 was adjusted for model 1 but participants with CVD, cancer, diabetes hypertension and major illness were excluded from the analyses.

Table S8. Association between genetic profile risk score and BMI levels of physical activity and sedentary behaviour

|  | n | Low levels |  | High levels |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | Beta (95\% CI) | p -value | Beta (95\% CI) | p -value | $\mathbf{P}_{\text {(interaction }}$ |
| Self-reported total PA | 338,216 | 0.63 (0.61 to 0.66) | $1.6 \times 10^{-7}$ | 0.46 (0.44 to 0.48) | $8.0 \times 10^{-10}$ | $3.8 \times 10^{-29}$ |
| Objective Total PA* | 62,881 | 0.55 (0.50 to 0.60) | $6.5 \times 10^{-14}$ | 0.40 (0.36 to 0.45) | $1.1 \times 10^{-83}$ | $3.5 \times 10^{-6}$ |
| Total discretionary sedentary time | 338,216 | 0.49 (0.47 to 0.51) | $3.8 \times 10^{-13}$ | 0.62 (0.60 to 0.65) | $4.0 \times 10^{-5}$ | $2.9 \times 10^{-17}$ |
| TV-viewing | 338,216 | 0.50 (0.48 to 0.52) | $6.3 \times 10^{-10}$ | 0.63 (0.60 to 0.66) | $2.1 \times 10^{-7}$ | $1.2 \times 10^{-13}$ |

Data presented as beta coefficients $(95 \% \mathrm{CI})$. The beta coefficient indicates the change in BMI per SD increase in the genetic profile risk score by high or low levels of physical activity (self-reported or objectively-measured PA) and high and low levels of discretionary sedentary behaviors (overall discretionary sedentary time and TV-viewing). The p-value for the interaction between GPRS and the exposure of interest (PA self-reported, PA accelerometer, sedentary behaviors and TV-viewing) is presented as Pinteraction. Analyses were adjusted for age, sex, deprivation, education qualifications, recruitment center, month of recruitment, the first 10 principal components of ancestry and genotyping batch, smoking status, dietary intake (alcohol, fruit \& vegetable, red meat, processed meat, cereals, bread and cheese) and comorbidities (diabetes, hypertension, cardiovascular diseases and cancer). Analyses performed for objectively measured PA were additionally adjusted for season and wearing time and sedentary behavior whereas analyses for sedentary behavior and TV-viewing were additionally adjusted for self-reported total PA.
*Objectively measured physical activity; PA: physical activity; BMI: body mass index.

Table S9. Association between genetic profile risk score and BMI by the combined categories of physical activity and sedentary behaviour

|  | n | Physically active \& Low sedentary behaviour |  | Physically active \& High sedentary behaviour |  | Physically inactive \& Low sedentary behaviour |  | Physically inactive \& High sedentary behaviour |  | P (interaction) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{CI}) \end{gathered}$ | p -value | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{Cl}) \end{gathered}$ | $p$-value | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{CI}) \end{gathered}$ | $p$-value | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{Cl}) \end{gathered}$ | $p$-value |  |
| Self-reported Total PA | 338,216 | $\begin{gathered} 0.43 \\ (0.40 \text { to } 0.46) \end{gathered}$ | $1.7 \times 10^{-279}$ | $\begin{gathered} 0.51 \\ (0.48 \text { to } 0.55) \\ \hline \end{gathered}$ | $8.4 \times 10^{-207}$ | $\begin{gathered} 0.55 \\ (0.51 \text { to } 0.59) \end{gathered}$ | $3.7 \times 10^{-250}$ | $\begin{gathered} 0.73 \\ (0.69 \text { to } 0.78) \end{gathered}$ | $1.3 \times 10^{-281}$ | $2.3 \times 10^{-34}$ |
| Objective Total PA* | 62,881 | $\begin{gathered} 0.37 \\ (0.32 \text { to } 0.42) \\ \hline \end{gathered}$ | $2.3 \times 10^{-51}$ | $\begin{gathered} 0.47 \\ (0.39 \text { to } 0.54) \\ \hline \end{gathered}$ | $5.6 \times 10^{-34}$ | $\begin{gathered} 0.52 \\ (0.45 \text { to } 0.58) \\ \hline \end{gathered}$ | $1.2 \times 10^{-54}$ | $\begin{gathered} 0.60 \\ (0.52 \text { to } 0.68) \\ \hline \end{gathered}$ | $8.9 \times 10^{-50}$ | $3.5 \times 10^{-6}$ |

Data presented as beta coefficients $(95 \% \mathrm{CI})$. The beta coefficient indicates the change in BMI per SD increase in the genetic profile risk score by combined categories of high and low physical activity (self-reported or objectively-measured PA) and overall discretionary sedentary behaviors. A multiplicative interaction term between GPRS-BMI and combined categories of PA/Sedentary behaviors were fitted into the model to investigate a potential interaction effect. The $p$-value for the interaction between GPRS and the exposure of interest (PA self-reported, PA accelerometer, sedentary behaviors and TV-viewing) is presented as P-interaction.
Analyses were adjusted for age, sex, deprivation, education qualifications, recruitment center, month of recruitment, the first 10 principal components of ancestry and genotyping batch, smoking status, dietary intake (alcohol, fruit \& vegetable, red meat, processed meat, cereals, bread and cheese) and comorbidities (diabetes, hypertension, cardiovascular diseases and cancer). Analyses performed for objectively measured PA were additionally adjusted for season and wearing time.
*objectively measured physical activity; PA: physical activity; BMI: body mass index.

Table S10. Association between genetic profile risk score and BMI by the combined categories of physical activity and TV-viewing

|  | n | Physically active \& Low TV-viewing |  | Physically active \& High TV-viewing |  | Physically inactive \& Low TV-viewing |  | Physically inactive \& High TV-viewing |  | $\mathbf{P}$ (interaction) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{CI}) \end{gathered}$ | $p$-value | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{Cl}) \end{gathered}$ | $p$-value | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{CI}) \end{gathered}$ | $p$-value | $\begin{gathered} \text { Beta } \\ (95 \% \mathrm{CI}) \end{gathered}$ | $p$-value |  |
| Self-reported Total PA | 338,216 | $\begin{gathered} 0.43 \\ (0.40 \text { to } 0.46) \end{gathered}$ | $2.0 \times 10^{-276}$ | $\begin{gathered} 0.52 \\ (0.48 \text { to } 0.56) \\ \hline \end{gathered}$ | $5.6 \times 10^{-163}$ | $\begin{gathered} 0.58 \\ (0.55 \text { to } 0.61) \\ \hline \end{gathered}$ | $2.8 \times 10^{-301}$ | $\begin{gathered} 0.72 \\ (0.68 \text { to } 0.76) \end{gathered}$ | $4.2 \times 10^{-251}$ | $4.4 \times 10^{-30}$ |
| Objective Total PA* | 62,774 | $\begin{gathered} 0.40 \\ (0.35 \text { to } 0.45) \\ \hline \end{gathered}$ | $5.1 \times 10^{-64}$ | $\begin{gathered} 0.44 \\ (0.36 \text { to } 0.54) \\ \hline \end{gathered}$ | $2.3 \times 10^{-22}$ | $\begin{gathered} 0.53 \\ (0.47 \text { to } 0.59) \\ \hline \end{gathered}$ | $2.6 \times 10^{-68}$ | $\begin{gathered} 0.60 \\ (0.51 \text { to } 0.70) \\ \hline \end{gathered}$ | $1.7 \times 10^{-35}$ | $2.9 \times 10^{-6}$ |

Data presented as beta coefficients ( $95 \% \mathrm{CI}$ ). The beta coefficient indicates the change in BMI per SD increase in the genetic profile risk score by combined categories of high and low physical activity (self-reported or objectively-measured PA) and overall discretionary sedentary behaviors. A multiplicative interaction term between GPRS-BMI and combined categories of PA/Sedentary behaviors were fitted into the model to investigate a potential interaction effect. The p-value for the interaction between GPRS and the exposure of interest (PA self-reported, PA accelerometer, sedentary behaviors and TV-viewing) is presented as P-interaction.
Analyses were adjusted for age, sex, deprivation, education qualifications, recruitment center, month of recruitment, the first 10 principal components of ancestry and genotyping batch, smoking status, dietary intake (alcohol, fruit \& vegetable, red meat, processed meat, cereals, bread and cheese) and comorbidities (diabetes, hypertension, cardiovascular diseases and cancer). Analyses performed for objectively measured PA were additionally adjusted for season and wearing time.
*objectively measured physical activity; PA: physical activity; BMI: body mass index.


[^0]:    ${ }^{1}$ Not genotyped in UK Biobank cohort and therefore not analysed in the current report.
    ${ }^{2}$ Excluded from the SNP set for GPRS calculation on the basis of Hardy-Weinberg equilibrium $p<10^{-6}$.

