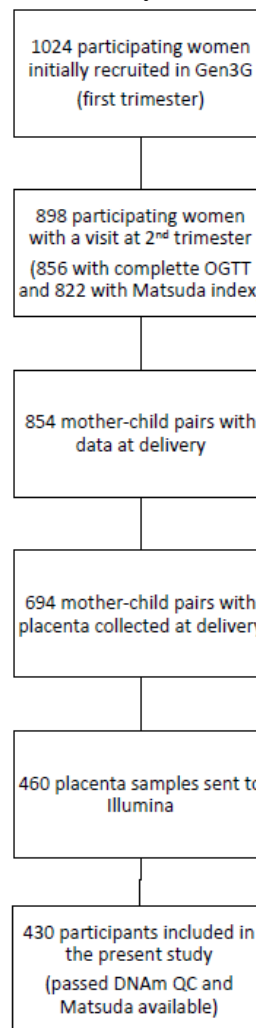
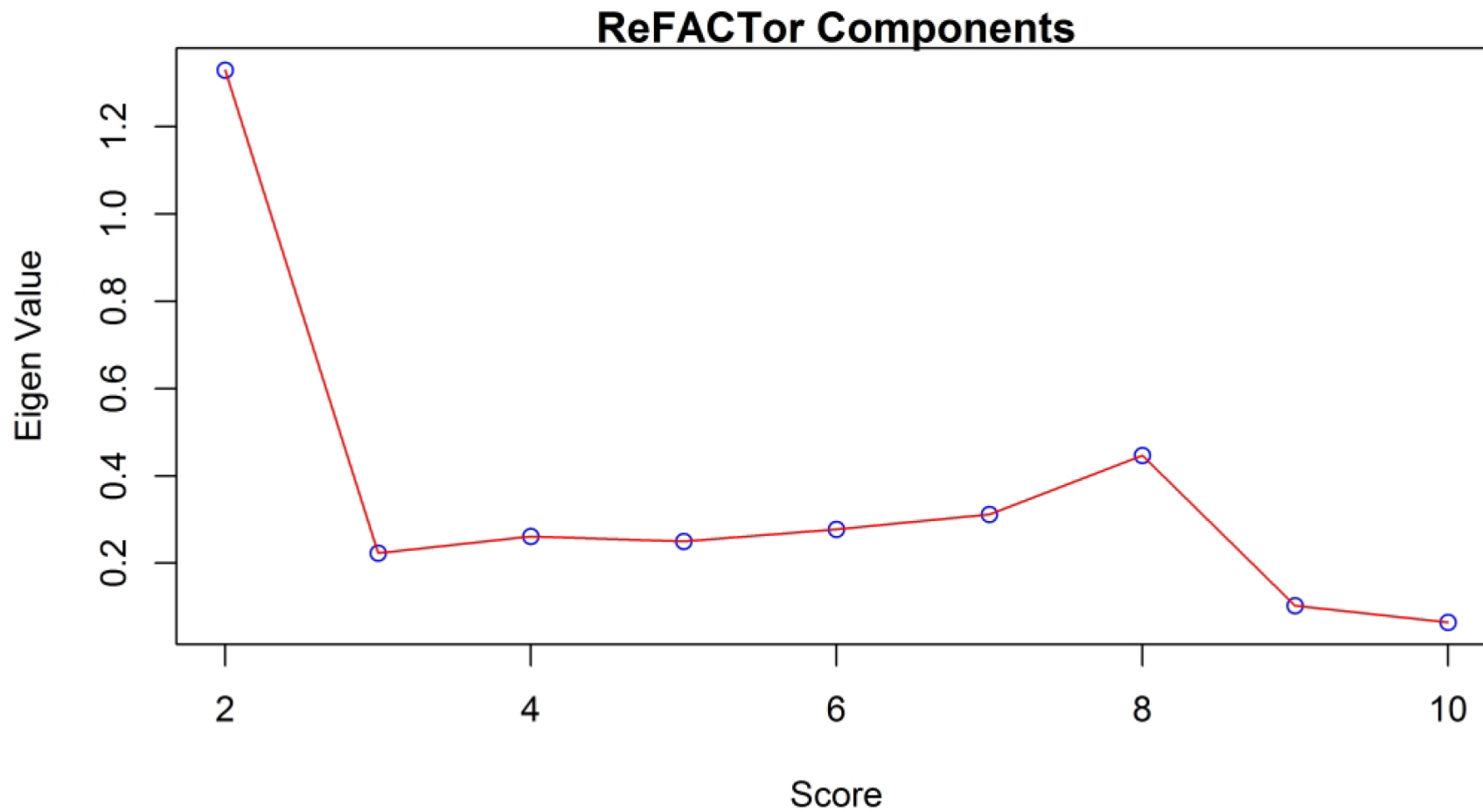


SUPPLEMENTARY DATA

Supplementary Figure 1. Participants flow chart included in current study

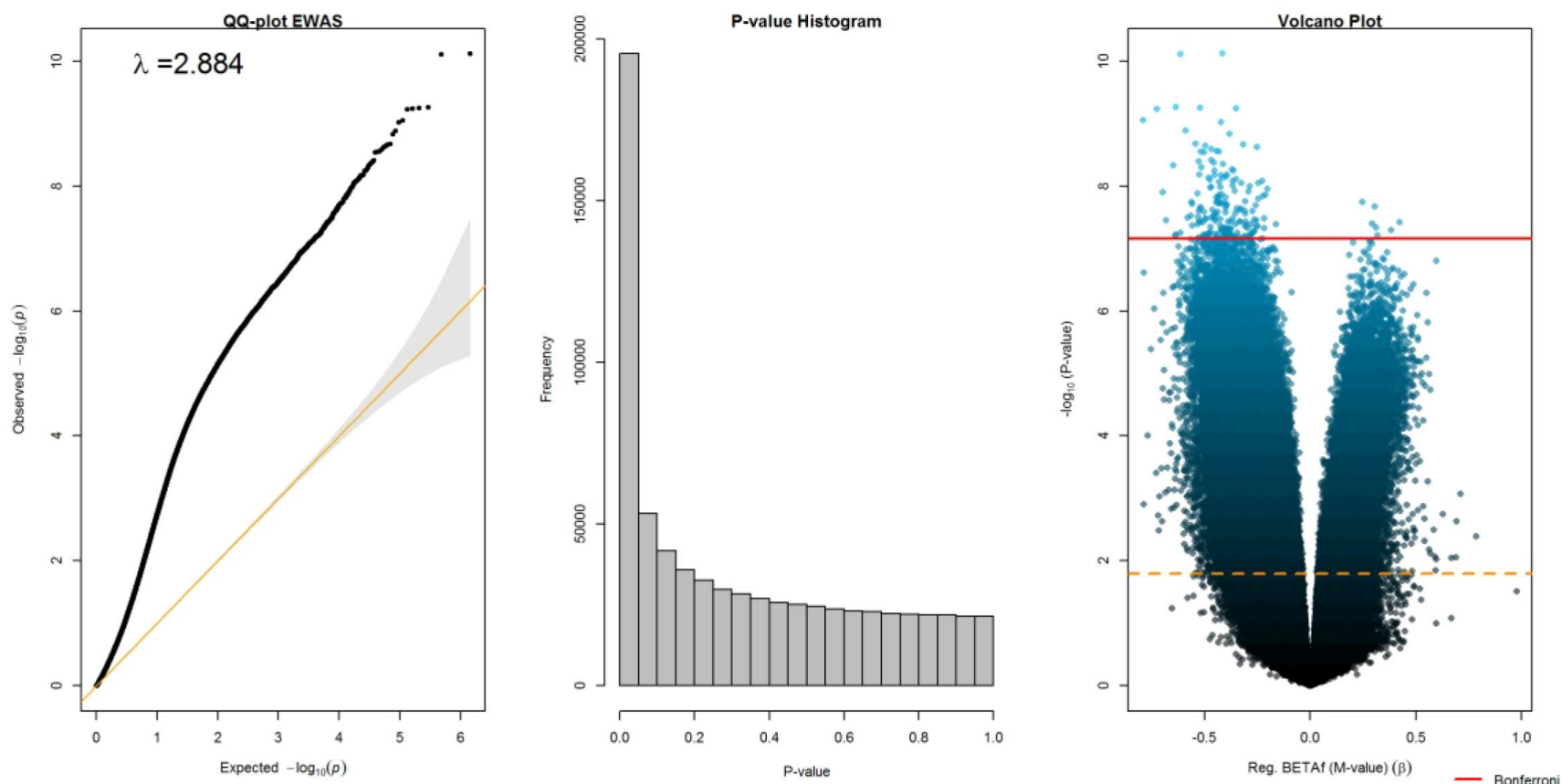


Supplementary Figure 2.

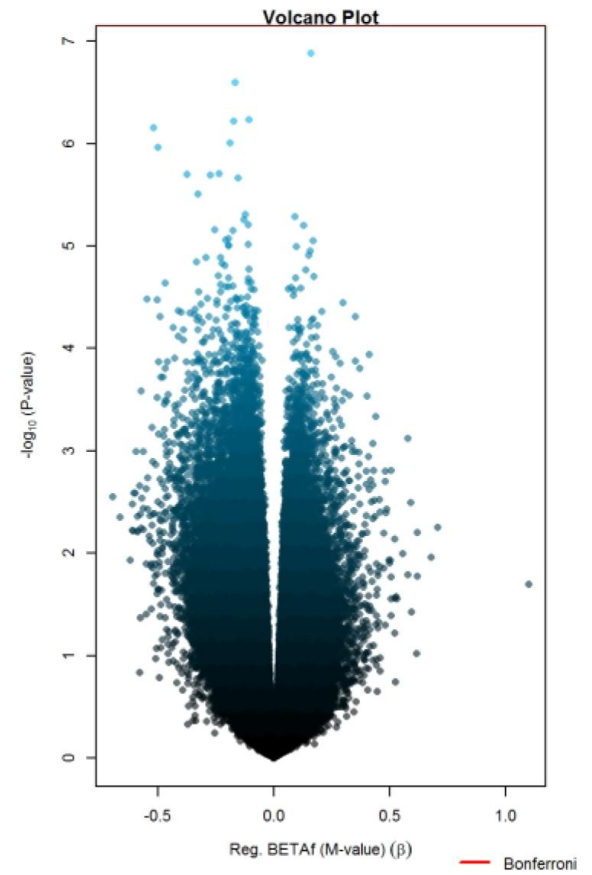
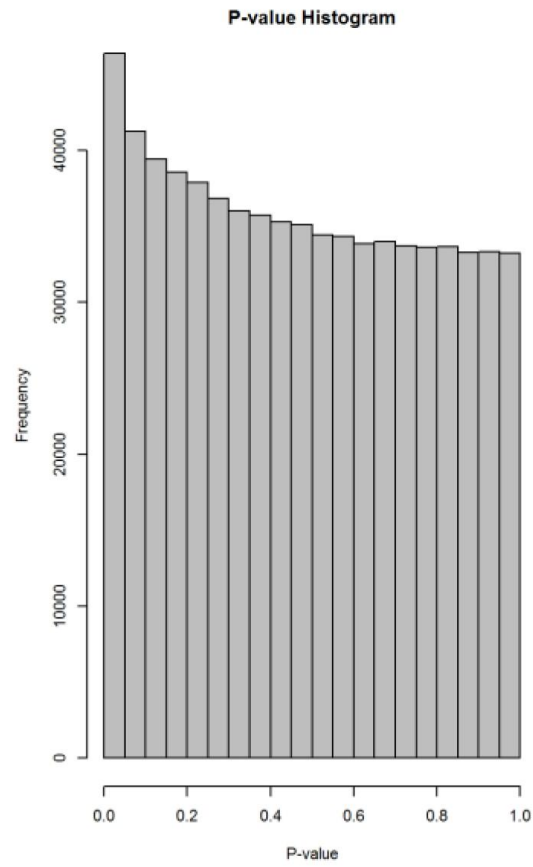
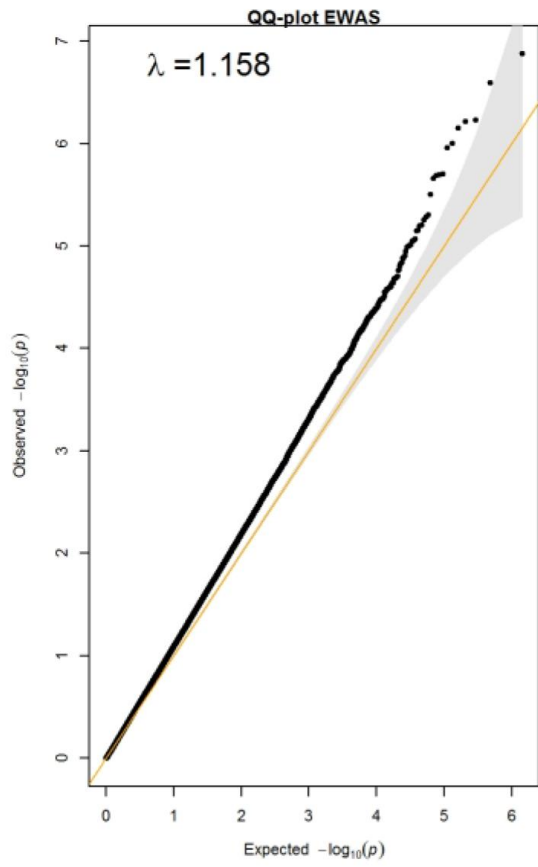


SUPPLEMENTARY DATA

Supplementary Figure 3.



SUPPLEMENTARY DATA



SUPPLEMENTARY DATA

| Supplementary Table 1: Single Nucleotide Polymorphisms (SNPs) used to create genetic risk score (GRS) of insulin sensitivity | | | | |
|--|--------------|-----------------------|---|--|
| SNPs | chr position | annotated gene | initial association in Prokenpenko et al. | Association in Gen3G with Matsuda (ln) |
| | | | Beta | beta (SE) |
| rs1260326 | 2:27730940 | <i>GCKR</i> | -0.067 | -0.02307 (0.03824) |
| rs35767 | 12:102875569 | <i>IGF1</i> | -0.080 | 0.01581 (0.05446) |
| rs459193 | 5:55806751 | <i>ANKRD55/MAP3K1</i> | -0.053 | -0.09932 (0.04122) |
| rs4846565 | 1:219722104 | <i>LYPLAL1</i> | -0.044 | -0.09431 (0.04130) |
| rs4865796 | 5:53272664 | <i>ARL15</i> | -0.071 | -0.01121 (0.04271) |
| rs6822892 | 4:157734675 | <i>PDGFC</i> | -0.093 | -0.006239 (0.040982) |
| rs7607980 | 2:165551201 | <i>COBLL1</i> | -0.120 | -0.09199 (0.05452) |
| rs1801282 | 3:12393125 | <i>PPARG</i> | -0.084 | -0.10657 (0.06494) |

SUPPLEMENTARY DATA

Supplementary Table 3. ReFACTor PCs and associations with Mastuda index and co-variables

| | Pval_PC1 | Pval_PC2 | Pval_PC3 | Pval_PC4 | Pval_PC5 | Pval_PC6 | Pval_PC7 | Pval_PC8 | Pval_PC9 | Pval_PC10 |
|--------------------------|----------|----------|----------|----------|----------|----------|----------|----------|----------|-----------|
| Matsuda index | <0.0001 | 0.011 | 0.495 | 0.123 | 0.021 | 0.068 | 0.603 | 0.822 | 0.505 | 0.932 |
| Maternal age | 0.202 | 0.080 | 0.715 | 0.642 | 0.726 | 0.034 | 0.225 | 0.614 | 0.432 | 0.163 |
| Maternal BMI (V1) | 0.314 | 0.700 | 0.407 | 0.570 | 0.756 | 0.433 | 0.801 | 0.367 | 0.971 | 0.193 |
| Gravid status | 0.635 | 0.338 | 0.289 | 0.351 | 0.598 | 0.212 | 0.146 | 0.350 | 0.299 | 0.307 |
| Maternal Smoking | 0.539 | 0.828 | 0.007 | 0.342 | 0.459 | 0.664 | 0.227 | 0.148 | 0.109 | 0.257 |
| Gestational age at birth | 0.289 | 0.736 | <0.0001 | 0.344 | 0.090 | 0.585 | 0.003 | 0.180 | 0.744 | 0.694 |
| Child sex | 0.834 | 0.578 | 0.469 | 0.073 | 0.398 | 0.140 | 0.580 | 0.883 | 0.441 | 0.007 |

P-values from correlations if continuous variables, and from Kruskal-Wallis if categorical variables

SUPPLEMENTARY DATA

Supplementary Table 4. Results from Mendelian Randomization analyses, building genetic instrumental variables (GRS or single SNP) for DNA methylation at 188 CpGs identified in Model 1 using cis-- - SNPs within 500kb around each CpG site

SUPPLEMENTARY DATA

| | | | GRS information on association with CpG and elastic net statistics (N = 418) | | | | | | | | | | | linear regression ln(matsuda) ~ GRS (N = 401) | | | | TSLS results (N = 401) | | | linear regression ln(matsuda) ~ CpG (N = 401) | | |
|------------|-----|-----------|--|-----------|------------|----------|-------|-------------------------|--------------------|--------------------------|------------------|----------|-------|---|---------|----------------------------------|----------|------------------------|---------|----------------------------------|---|-------|----------|
| CPG | chr | pos | Estimate | Std.Error | P.Value | r.square | Fstat | N of SNPs within ±500kb | N of SNPs MAF<0.05 | N SNPs without duplicate | N of SNPs in GRS | estimate | SE | P_value | Rsquare | P_value adj for multiple testing | estimate | SE | P_value | P_value adj for multiple testing | estimate | SE | P_value |
| cg12658750 | 18 | 75499382 | 0.169 | 0.028 | 3.37E-09 | 0.081 | 36.5 | 16935 | 3291 | 1811 | 5 | -0.079 | 0.023 | 0.0008 | 0.028 | 0.03 | -0.448 | 0.142 | 0.0017 | 0.046 | -0.150 | 0.039 | 1.45E-04 |
| cg24475484 | 8 | 1625395 | 0.189 | 0.025 | 2.81E-13 | 0.120 | 57.0 | 25101 | 4134 | 2769 | 10 | -0.046 | 0.014 | 0.0009 | 0.027 | 0.03 | -0.242 | 0.074 | 0.0012 | 0.046 | -0.103 | 0.025 | 5.09E-05 |
| cg01618245 | 20 | 61990279 | 0.113 | 0.012 | 3.20E-20 | 0.185 | 94.3 | 18168 | 2970 | 1608 | 34 | -0.022 | 0.007 | 0.0010 | 0.027 | 0.03 | -0.195 | 0.059 | 0.0011 | 0.046 | -0.106 | 0.025 | 3.31E-05 |
| cg08099672 | 9 | 139944162 | 0.095 | 0.013 | 2.76E-12 | 0.111 | 51.9 | 17133 | 1692 | 1077 | 13 | -0.034 | 0.010 | 0.0012 | 0.026 | 0.03 | -0.342 | 0.107 | 0.0015 | 0.046 | -0.156 | 0.036 | 1.97E-05 |
| cg12673377 | 7 | 1388271 | 0.062 | 0.005 | 2.37E-27 | 0.246 | 135.8 | 20418 | 2694 | 1672 | 51 | -0.015 | 0.004 | 0.0012 | 0.026 | 0.03 | -0.238 | 0.073 | 0.0012 | 0.046 | -0.150 | 0.036 | 3.07E-05 |
| cg03699074 | 16 | 88849875 | 0.095 | 0.014 | 2.21E-11 | 0.102 | 47.3 | 21611 | 2782 | 1905 | 17 | 0.022 | 0.007 | 0.0030 | 0.022 | 0.07 | 0.224 | 0.078 | 0.0044 | 0.10 | 0.075 | 0.025 | 3.13E-03 |
| cg24073146 | 11 | 2543204 | 0.108 | 0.017 | 1.35E-09 | 0.085 | 38.5 | 17447 | 2473 | 1698 | 6 | -0.046 | 0.016 | 0.0039 | 0.021 | 0.07 | -0.416 | 0.150 | 0.006 | 0.11 | -0.140 | 0.042 | 8.89E-04 |
| cg08511578 | 10 | 1403938 | 0.106 | 0.024 | 1.50E-05 | 0.044 | 19.2 | 17754 | 2647 | 1657 | 4 | -0.034 | 0.013 | 0.0079 | 0.018 | 0.13 | -0.327 | 0.134 | 0.015 | 0.16 | -0.098 | 0.025 | 8.29E-05 |
| cg11765907 | 20 | 43103949 | 0.047 | 0.005 | 1.50E-16 | 0.151 | 74.2 | 14001 | 1783 | 1041 | 27 | -0.016 | 0.006 | 0.010 | 0.016 | 0.15 | -0.324 | 0.125 | 0.010 | 0.16 | -0.187 | 0.050 | 1.95E-04 |
| cg02384870 | 2 | 216161208 | 0.175 | 0.033 | 1.53E-07 | 0.064 | 28.5 | 14134 | 1835 | 1063 | 5 | -0.042 | 0.017 | 0.014 | 0.015 | 0.17 | -0.243 | 0.100 | 0.016 | 0.16 | -0.112 | 0.024 | 4.84E-06 |
| cg21396983 | 16 | 1275423 | 0.079 | 0.009 | 1.39E-16 | 0.152 | 74.3 | 21550 | 2710 | 1738 | 24 | -0.015 | 0.006 | 0.016 | 0.015 | 0.17 | -0.182 | 0.074 | 0.014 | 0.16 | -0.144 | 0.030 | 1.97E-06 |
| cg09640571 | 16 | 88388040 | 0.031 | 0.004 | 1.75E-12 | 0.113 | 52.9 | 21289 | 3220 | 2169 | 27 | -0.011 | 0.005 | 0.016 | 0.014 | 0.17 | -0.356 | 0.147 | 0.016 | 0.16 | -0.211 | 0.049 | 1.71E-05 |
| cg24857040 | 13 | 113565214 | 0.079 | 0.008 | 5.26E-21 | 0.192 | 98.7 | 17276 | 2194 | 1324 | 26 | -0.013 | 0.006 | 0.018 | 0.014 | 0.17 | -0.169 | 0.071 | 0.017 | 0.16 | -0.126 | 0.032 | 7.73E-05 |
| cg20612875 | 17 | 1094308 | 0.035 | 0.003 | 4.26E-22 | 0.201 | 104.9 | 18826 | 2792 | 1917 | 39 | -0.009 | 0.004 | 0.020 | 0.014 | 0.17 | -0.252 | 0.106 | 0.017 | 0.16 | -0.224 | 0.049 | 7.78E-06 |
| cg23521973 | 10 | 125117429 | 0.079 | 0.011 | 7.45E-13 | 0.116 | 54.8 | 15015 | 2399 | 1336 | 17 | -0.017 | 0.007 | 0.020 | 0.014 | 0.17 | -0.212 | 0.090 | 0.019 | 0.16 | -0.138 | 0.031 | 1.13E-05 |
| cg14440140 | 9 | 140177567 | 0.081 | 0.008 | 8.24E-21 | 0.190 | 97.6 | 16195 | 1470 | 921 | 34 | -0.013 | 0.006 | 0.025 | 0.012 | 0.19 | -0.154 | 0.068 | 0.024 | 0.19 | -0.109 | 0.030 | 3.10E-04 |
| cg06429120 | 16 | 86309214 | 0.155 | 0.013 | 9.85E-28 | 0.249 | 138.1 | 22860 | 3698 | 2812 | 36 | -0.015 | 0.007 | 0.027 | 0.012 | 0.19 | -0.097 | 0.043 | 0.023 | 0.19 | -0.101 | 0.021 | 1.86E-06 |
| cg00460471 | 2 | 61437605 | 0.102 | 0.024 | 3.10E-05 | 0.041 | 17.7 | 13349 | 1511 | 814 | 3 | -0.032 | 0.014 | 0.028 | 0.012 | 0.19 | -0.319 | 0.151 | 0.035 | 0.21 | -0.118 | 0.028 | 3.55E-05 |
| cg02110836 | 17 | 71953083 | 0.057 | 0.005 | 2.91E-24 | 0.220 | 117.4 | 17019 | 2437 | 1629 | 30 | -0.015 | 0.007 | 0.028 | 0.012 | 0.19 | -0.265 | 0.119 | 0.026 | 0.19 | -0.227 | 0.056 | 5.57E-05 |
| cg06077821 | 5 | 180048893 | 0.047 | 0.005 | 2.68E-19 | 0.176 | 89.1 | 16657 | 2554 | 1497 | 34 | -0.008 | 0.004 | 0.031 | 0.012 | 0.20 | -0.178 | 0.081 | 0.028 | 0.19 | -0.146 | 0.034 | 2.43E-05 |
| cg00875983 | 11 | 3240213 | 0.071 | 0.008 | 1.34E-18 | 0.170 | 85.3 | 17171 | 2667 | 1744 | 20 | -0.017 | 0.008 | 0.033 | 0.011 | 0.21 | -0.237 | 0.109 | 0.031 | 0.20 | -0.176 | 0.045 | 1.16E-04 |
| cg14361947 | 9 | 140780551 | 0.056 | 0.007 | 5.95E-15 | 0.136 | 65.7 | 12059 | 978 | 679 | 24 | -0.016 | 0.008 | 0.041 | 0.010 | 0.22 | -0.295 | 0.144 | 0.041 | 0.21 | -0.161 | 0.052 | 2.02E-03 |
| cg00386725 | 1 | 18617791 | 0.095 | 0.012 | 6.12E-15 | 0.136 | 65.6 | 16267 | 2471 | 1494 | 12 | -0.024 | 0.012 | 0.041 | 0.010 | 0.22 | -0.248 | 0.120 | 0.038 | 0.21 | -0.185 | 0.044 | 3.53E-05 |
| cg10964639 | 16 | 24043296 | 0.059 | 0.008 | 4.62E-13 | 0.118 | 55.9 | 14736 | 2182 | 1222 | 24 | -0.011 | 0.005 | 0.041 | 0.010 | 0.22 | -0.182 | 0.088 | 0.039 | 0.21 | -0.122 | 0.031 | 9.72E-05 |
| cg11855710 | 19 | 29768180 | 0.148 | 0.021 | 1.85E-11 | 0.103 | 47.7 | 17654 | 3015 | 1361 | 10 | -0.029 | 0.014 | 0.043 | 0.010 | 0.22 | -0.199 | 0.096 | 0.039 | 0.21 | -0.150 | 0.031 | 2.42E-06 |
| cg12954267 | 6 | 33093109 | 0.092 | 0.026 | 4.69E-04 | 0.029 | 12.4 | 18517 | 6412 | 2328 | 3 | -0.028 | 0.014 | 0.045 | 0.010 | 0.22 | -0.321 | 0.174 | 0.07 | 0.27 | -0.089 | 0.025 | 5.30E-04 |
| cg00389386 | 1 | 6188136 | 0.062 | 0.007 | 8.31E-17 | 0.154 | 75.5 | 16477 | 1981 | 1136 | 22 | -0.014 | 0.007 | 0.045 | 0.010 | 0.22 | -0.234 | 0.115 | 0.043 | 0.21 | -0.174 | 0.044 | 1.06E-04 |
| cg21347874 | 17 | 77122733 | 0.083 | 0.012 | 6.99E-11 | 0.097 | 44.8 | 16022 | 2400 | 1721 | 13 | -0.016 | 0.008 | 0.047 | 0.010 | 0.22 | -0.188 | 0.093 | 0.044 | 0.21 | -0.127 | 0.029 | 1.26E-05 |
| cg27625886 | 20 | 62060443 | 0.135 | 0.015 | 8.29E-17 | 0.154 | 75.6 | 18081 | 2937 | 1609 | 17 | -0.018 | 0.009 | 0.05 | 0.009 | 0.23 | -0.128 | 0.064 | 0.048 | 0.23 | -0.108 | 0.026 | 3.90E-05 |
| cg11586089 | 14 | 66644923 | 0.090 | 0.010 | 2.11E-17 | 0.159 | 78.8 | 13997 | 1665 | 708 | 35 | -0.013 | 0.007 | 0.06 | 0.009 | 0.25 | -0.147 | 0.077 | 0.06 | 0.25 | -0.110 | 0.030 | 2.96E-04 |
| cg16340422 | 17 | 17110120 | 0.137 | 0.020 | 6.55E-11 | 0.098 | 45.0 | 13630 | 1744 | 1022 | 10 | -0.025 | 0.013 | 0.06 | 0.009 | 0.26 | -0.187 | 0.098 | 0.06 | 0.25 | -0.124 | 0.029 | 2.89E-05 |
| cg18861231 | 3 | 152057943 | 0.035 | 0.007 | 3.48E-07 | 0.061 | 26.8 | 13018 | 1777 | 676 | 18 | -0.007 | 0.004 | 0.06 | 0.009 | 0.26 | -0.195 | 0.106 | 0.07 | 0.27 | -0.079 | 0.025 | 1.53E-03 |
| cg03568769 | 11 | 1800837 | 0.074 | 0.005 | 1.28E-36 | 0.319 | 195.0 | 17582 | 2953 | 1759 | 66 | -0.008 | 0.004 | 0.07 | 0.008 | 0.27 | -0.104 | 0.057 | 0.07 | 0.27 | -0.117 | 0.032 | 3.12E-04 |
| cg12760575 | 9 | 139158654 | 0.110 | 0.013 | 5.14E-16 | 0.146 | 71.3 | 17109 | 1972 | 1300 | 15 | -0.020 | 0.011 | 0.07 | 0.008 | 0.28 | -0.185 | 0.102 | 0.07 | 0.27 | -0.151 | 0.039 | 1.47E-04 |
| cg26037026 | 7 | 1641922 | 0.194 | 0.055 | 0.000489 | 0.029 | 12.4 | 20490 | 2519 | 1493 | 1 | -0.078 | 0.045 | 0.08 | 0.008 | 0.30 | -0.406 | 0.241 | 0.09 | 0.30 | -0.143 | 0.039 | 2.39E-04 |
| cg14839558 | 1 | 1874488 | 0.141 | 0.017 | 3.66E-15 | 0.138 | 66.8 | 17112 | 1789 | 1113 | 19 | -0.018 | 0.010 | 0.08 | 0.008 | 0.30 | -0.133 | 0.075 | 0.08 | 0.29 | -0.104 | 0.027 | 1.38E-04 |
| cg24600306 | 20 | 61990671 | 0.070 | 0.007 | 5.23E-19 | 0.174 | 87.5 | 18172 | 2970 | 1608 | 24 | -0.013 | 0.008 | 0.08 | 0.007 | 0.30 | -0.193 | 0.110 | 0.08 | 0.29 | -0.174 | 0.046 | 1.91E-04 |
| cg16853336 | 3 | 149396490 | 0.040 | 0.006 | 6.82E-11 | 0.097 | 44.9 | 13981 | 2115 | 1079 | 19 | -0.011 | 0.006 | 0.09 | 0.007 | 0.30 | -0.270 | 0.157 | 0.09 | 0.30 | -0.174 | 0.050 | 5.04E-04 |
| cg08149223 | 10 | 50507130 | 0.043 | 0.007 | 1.56E-09 | 0.084 | 38.1 | 15782 | 2177 | 1029 | 16 | -0.012 | 0.007 | 0.09 | 0.007 | 0.30 | -0.273 | 0.159 | 0.09 | 0.30 | -0.171 | 0.047 | 2.86E-04 |
| cg00781180 | 16 | 73984002 | 0.218 | 0.037 | 4.82E-09 | 0.079 | 35.8 | 15775 | 1655 | 1261 | 6 | -0.038 | 0.023 | 0.09 | 0.007 | 0.30 | -0.177 | 0.103 | 0.09 | 0.30 | -0.111 | 0.028 | 1.11E-04 |
| cg16444062 | 16 | 2770298 | 0.236 | 0.053 | 1.03E-05 | 0.046 | 19.9 | 15567 | 1480 | 949 | 3 | -0.047 | 0.029 | 0.10 | 0.007 | 0.32 | -0.192 | 0.117 | 0.10 | 0.31 | -0.105 | 0.026 | 4.92E-05 |
| cg24738497 | 19 | 36005395 | 0.130 | 0.024 | 5.27E-08 | 0.069 | 30.7 | 14689 | 2074 | 1243 | 4 | -0.028 | 0.017 | 0.10 | 0.007 | 0.32 | -0.206 | 0.124 | 0.10 | 0.31 | -0.149 | 0.034 | 1.20E-05 |
| cg17756860 | 4 | 110510374 | 0.122 | 0.034 | 0.00034286 | 0.030 | 13.0 | 13975 | 1499 | 674 | 4 | -0.032 | 0.020 | 0.11 | 0.007 | 0.32 | -0.264 | 0.163 | 0.11 | 0.31 | -0.128 | 0.028 | 6.14E-06 |
| cg08737919 | 12 | 108840669 | 0.207 | 0.034 | 2.47E-09 | 0.082 | 37.2 | 14629 | 2021 | 914 | 6 | -0.034 | 0.021 | 0.11 | 0.006 | 0.32 | -0.163 | 0.101 | 0.11 | 0.31 | -0.095 | 0.030 | 1.37E-03 |

SUPPLEMENTARY DATA

| | | | | | | | | | | | | | | | | | | | | | | | |
|------------|----|-----------|-------|-------|------------|-------|-------|-------|------|------|----|--------|-------|------|-------|------|--------|-------|------|------|--------|-------|----------|
| cg12193237 | 19 | 3907913 | 0.048 | 0.005 | 8.51E-18 | 0.163 | 80.9 | 16921 | 2133 | 1364 | 36 | -0.007 | 0.004 | 0.11 | 0.006 | 0.32 | -0.147 | 0.090 | 0.10 | 0.31 | -0.143 | 0.036 | 8.65E-05 |
| cg07435932 | 13 | 19648633 | 0.381 | 0.127 | 0.00280574 | 0.021 | 9.0 | 16939 | 2652 | 1044 | 1 | -0.102 | 0.065 | 0.12 | 0.006 | 0.34 | -0.283 | 0.192 | 0.14 | 0.36 | -0.091 | 0.025 | 2.94E-04 |
| cg13449257 | 9 | 137355128 | 0.055 | 0.009 | 1.52E-08 | 0.074 | 33.3 | 18196 | 2544 | 1691 | 14 | -0.011 | 0.007 | 0.13 | 0.006 | 0.36 | -0.191 | 0.125 | 0.13 | 0.36 | -0.145 | 0.034 | 2.35E-05 |
| cg02287371 | 5 | 1246274 | 0.157 | 0.037 | 2.90E-05 | 0.041 | 17.9 | 17997 | 2289 | 1564 | 3 | -0.033 | 0.022 | 0.13 | 0.006 | 0.36 | -0.208 | 0.139 | 0.14 | 0.36 | -0.097 | 0.028 | 4.76E-04 |
| cg06751809 | 3 | 105047227 | 0.106 | 0.019 | 2.81E-08 | 0.072 | 32.0 | 14927 | 2475 | 943 | 13 | -0.012 | 0.008 | 0.13 | 0.006 | 0.36 | -0.115 | 0.075 | 0.13 | 0.36 | -0.086 | 0.020 | 3.58E-05 |
| cg01948226 | 6 | 32829208 | 0.092 | 0.015 | 1.24E-09 | 0.085 | 38.6 | 19534 | 8426 | 2926 | 10 | -0.014 | 0.010 | 0.14 | 0.005 | 0.37 | -0.159 | 0.106 | 0.13 | 0.36 | -0.127 | 0.030 | 2.87E-05 |
| cg00046877 | 22 | 19702664 | 0.065 | 0.009 | 2.83E-12 | 0.111 | 51.8 | 15502 | 2141 | 1221 | 12 | -0.016 | 0.011 | 0.15 | 0.005 | 0.38 | -0.248 | 0.168 | 0.14 | 0.36 | -0.209 | 0.057 | 2.84E-04 |
| cg01864178 | 20 | 44662890 | 0.087 | 0.014 | 1.25E-09 | 0.085 | 38.6 | 14114 | 1927 | 859 | 15 | -0.012 | 0.009 | 0.16 | 0.005 | 0.38 | -0.148 | 0.102 | 0.15 | 0.37 | -0.129 | 0.029 | 1.16E-05 |
| cg18682781 | 5 | 176277687 | 0.190 | 0.042 | 8.45E-06 | 0.047 | 20.3 | 14377 | 1242 | 712 | 3 | -0.035 | 0.025 | 0.16 | 0.005 | 0.38 | -0.181 | 0.127 | 0.16 | 0.37 | -0.117 | 0.028 | 3.03E-05 |
| cg13463516 | 9 | 138632121 | 0.035 | 0.005 | 5.61E-11 | 0.098 | 45.3 | 18674 | 2377 | 1670 | 24 | -0.006 | 0.004 | 0.17 | 0.005 | 0.38 | -0.163 | 0.115 | 0.16 | 0.37 | -0.147 | 0.036 | 5.41E-05 |
| cg09319487 | 20 | 56771178 | 0.148 | 0.012 | 4.59E-28 | 0.252 | 140.1 | 15943 | 2235 | 1345 | 28 | -0.012 | 0.009 | 0.17 | 0.005 | 0.38 | -0.083 | 0.060 | 0.16 | 0.37 | -0.101 | 0.030 | 9.53E-04 |
| cg08204714 | 16 | 87726461 | 0.201 | 0.064 | 0.00182003 | 0.023 | 9.8 | 21918 | 2969 | 2053 | 1 | -0.093 | 0.067 | 0.17 | 0.005 | 0.38 | -0.475 | 0.355 | 0.18 | 0.37 | -0.180 | 0.049 | 2.58E-04 |
| cg24064348 | 1 | 54181174 | 0.056 | 0.010 | 3.99E-08 | 0.070 | 31.3 | 13794 | 1697 | 881 | 8 | -0.017 | 0.012 | 0.17 | 0.005 | 0.38 | -0.310 | 0.224 | 0.17 | 0.37 | -0.215 | 0.056 | 1.65E-04 |
| cg18216439 | 18 | 35132882 | 0.209 | 0.065 | 0.0013874 | 0.024 | 10.4 | 13311 | 1742 | 916 | 1 | -0.055 | 0.041 | 0.18 | 0.004 | 0.38 | -0.280 | 0.212 | 0.19 | 0.37 | -0.128 | 0.030 | 2.65E-05 |
| cg04535454 | 8 | 144035192 | 0.049 | 0.009 | 2.17E-08 | 0.073 | 32.6 | 17157 | 2585 | 1321 | 16 | -0.009 | 0.006 | 0.18 | 0.004 | 0.38 | -0.175 | 0.129 | 0.18 | 0.37 | -0.134 | 0.034 | 1.02E-04 |
| cg08582801 | 7 | 99588335 | 0.041 | 0.009 | 6.73E-06 | 0.048 | 20.8 | 11898 | 913 | 363 | 7 | -0.014 | 0.011 | 0.19 | 0.004 | 0.38 | -0.332 | 0.247 | 0.18 | 0.37 | -0.232 | 0.056 | 3.89E-05 |
| cg27652925 | 6 | 31740951 | 0.049 | 0.010 | 1.90E-06 | 0.053 | 23.3 | 15655 | 5253 | 2035 | 8 | -0.015 | 0.011 | 0.19 | 0.004 | 0.38 | -0.307 | 0.230 | 0.18 | 0.37 | -0.198 | 0.052 | 1.73E-04 |
| cg09073467 | 1 | 6155515 | 0.073 | 0.006 | 1.34E-26 | 0.240 | 131.2 | 16527 | 2041 | 1179 | 36 | -0.009 | 0.007 | 0.19 | 0.004 | 0.38 | -0.126 | 0.095 | 0.18 | 0.37 | -0.185 | 0.046 | 8.07E-05 |
| cg08239663 | 16 | 52688569 | 0.210 | 0.044 | 2.29E-06 | 0.052 | 23.0 | 15429 | 2052 | 995 | 5 | -0.028 | 0.022 | 0.19 | 0.004 | 0.38 | -0.139 | 0.105 | 0.19 | 0.37 | -0.092 | 0.023 | 1.05E-04 |
| cg26225794 | 7 | 27717723 | 0.148 | 0.043 | 0.00074188 | 0.027 | 11.6 | 14250 | 1939 | 985 | 2 | 0.040 | 0.030 | 0.19 | 0.004 | 0.38 | 0.241 | 0.207 | 0.24 | 0.41 | -0.121 | 0.033 | 3.24E-04 |
| cg04293688 | 2 | 75466188 | 0.058 | 0.007 | 2.15E-14 | 0.131 | 62.8 | 15363 | 2334 | 1330 | 27 | -0.006 | 0.005 | 0.19 | 0.004 | 0.38 | -0.109 | 0.082 | 0.19 | 0.37 | -0.115 | 0.029 | 1.11E-04 |
| cg05202700 | 8 | 38258250 | 0.125 | 0.022 | 2.64E-08 | 0.072 | 32.2 | 12658 | 1326 | 661 | 8 | -0.016 | 0.013 | 0.19 | 0.004 | 0.38 | -0.125 | 0.095 | 0.19 | 0.37 | -0.103 | 0.026 | 1.15E-04 |
| cg15867652 | 17 | 79011277 | 0.119 | 0.012 | 9.22E-22 | 0.198 | 102.9 | 18385 | 2927 | 1635 | 23 | -0.009 | 0.007 | 0.20 | 0.004 | 0.38 | -0.075 | 0.057 | 0.19 | 0.37 | -0.104 | 0.026 | 6.38E-05 |
| cg00903007 | 17 | 34068171 | 0.020 | 0.002 | 2.67E-15 | 0.140 | 67.5 | 13632 | 1860 | 892 | 52 | -0.004 | 0.003 | 0.20 | 0.004 | 0.38 | -0.193 | 0.147 | 0.19 | 0.37 | -0.215 | 0.054 | 9.47E-05 |
| cg13377560 | 22 | 43860820 | 0.071 | 0.010 | 3.25E-12 | 0.110 | 51.5 | 16164 | 2593 | 1472 | 18 | -0.010 | 0.008 | 0.21 | 0.004 | 0.39 | -0.129 | 0.101 | 0.20 | 0.38 | -0.142 | 0.036 | 8.46E-05 |
| cg01095004 | 16 | 88454883 | 0.130 | 0.018 | 2.07E-12 | 0.112 | 52.5 | 21315 | 3131 | 2093 | 12 | -0.014 | 0.011 | 0.21 | 0.004 | 0.39 | -0.107 | 0.084 | 0.20 | 0.38 | -0.105 | 0.028 | 2.43E-04 |
| cg24848724 | 8 | 142578640 | 0.095 | 0.009 | 2.93E-21 | 0.194 | 100.1 | 16785 | 2282 | 1389 | 32 | -0.007 | 0.005 | 0.21 | 0.004 | 0.39 | -0.073 | 0.058 | 0.20 | 0.38 | -0.088 | 0.025 | 3.51E-04 |
| cg01898225 | 15 | 99975141 | 0.062 | 0.006 | 2.03E-19 | 0.178 | 89.8 | 15697 | 2792 | 1539 | 29 | -0.007 | 0.006 | 0.22 | 0.004 | 0.39 | -0.114 | 0.090 | 0.21 | 0.38 | -0.179 | 0.038 | 2.86E-06 |
| cg20018106 | 4 | 21768009 | 0.058 | 0.011 | 6.20E-08 | 0.068 | 30.4 | 16045 | 2280 | 975 | 13 | -0.007 | 0.006 | 0.22 | 0.004 | 0.39 | -0.123 | 0.097 | 0.21 | 0.38 | -0.121 | 0.026 | 5.69E-06 |
| cg18450810 | 17 | 51111066 | 0.214 | 0.045 | 3.11E-06 | 0.051 | 22.4 | 14484 | 2171 | 635 | 4 | -0.034 | 0.027 | 0.22 | 0.004 | 0.39 | -0.157 | 0.127 | 0.21 | 0.38 | -0.104 | 0.028 | 2.82E-04 |
| cg25285919 | 3 | 188312215 | 0.080 | 0.011 | 5.43E-13 | 0.118 | 55.5 | 14200 | 1596 | 1007 | 14 | -0.009 | 0.007 | 0.22 | 0.004 | 0.39 | -0.114 | 0.092 | 0.21 | 0.38 | -0.114 | 0.031 | 2.51E-04 |
| cg18184975 | 20 | 3644192 | 0.076 | 0.014 | 4.39E-08 | 0.070 | 31.1 | 14390 | 2558 | 1274 | 10 | -0.011 | 0.009 | 0.23 | 0.004 | 0.39 | -0.153 | 0.124 | 0.22 | 0.38 | -0.138 | 0.032 | 2.40E-05 |
| cg04131502 | 1 | 2845538 | 0.107 | 0.017 | 4.55E-10 | 0.089 | 40.8 | 17240 | 2079 | 1306 | 9 | -0.014 | 0.012 | 0.25 | 0.003 | 0.42 | -0.134 | 0.113 | 0.24 | 0.41 | -0.136 | 0.033 | 6.26E-05 |
| cg26849382 | 5 | 140901022 | 0.223 | 0.081 | 0.00651404 | 0.018 | 7.5 | 13286 | 1785 | 904 | 1 | 0.049 | 0.042 | 0.25 | 0.003 | 0.42 | 0.213 | 0.191 | 0.27 | 0.43 | 0.060 | 0.025 | 1.83E-02 |
| cg16819051 | 2 | 229173235 | 0.189 | 0.063 | 0.00303083 | 0.021 | 8.9 | 14534 | 2218 | 867 | 1 | -0.045 | 0.040 | 0.26 | 0.003 | 0.42 | -0.235 | 0.208 | 0.26 | 0.43 | -0.112 | 0.030 | 1.86E-04 |
| cg25721832 | 3 | 81913557 | 0.089 | 0.015 | 2.94E-09 | 0.081 | 36.8 | 12811 | 1396 | 484 | 17 | -0.012 | 0.010 | 0.27 | 0.003 | 0.43 | -0.132 | 0.118 | 0.26 | 0.43 | -0.101 | 0.033 | 2.13E-03 |
| cg22509138 | 20 | 44809026 | 0.357 | 0.119 | 0.00283299 | 0.021 | 9.0 | 14092 | 1752 | 957 | 1 | 0.070 | 0.064 | 0.27 | 0.003 | 0.43 | 0.194 | 0.201 | 0.33 | 0.50 | -0.107 | 0.026 | 3.62E-05 |
| cg05808230 | 20 | 61726456 | 0.095 | 0.011 | 1.17E-15 | 0.143 | 69.4 | 18272 | 2767 | 1736 | 19 | -0.009 | 0.008 | 0.27 | 0.003 | 0.43 | -0.092 | 0.083 | 0.27 | 0.43 | -0.131 | 0.032 | 4.96E-05 |
| cg09063835 | 20 | 58181392 | 0.152 | 0.020 | 2.26E-13 | 0.121 | 57.5 | 15305 | 1716 | 1164 | 13 | -0.013 | 0.012 | 0.28 | 0.003 | 0.43 | -0.088 | 0.079 | 0.27 | 0.43 | -0.117 | 0.027 | 1.78E-05 |
| cg06066032 | 13 | 48369175 | 0.370 | 0.116 | 0.00154941 | 0.024 | 10.2 | 14559 | 2328 | 815 | 1 | 0.061 | 0.059 | 0.30 | 0.003 | 0.47 | 0.185 | 0.203 | 0.36 | 0.53 | -0.098 | 0.025 | 7.18E-05 |
| cg05389237 | 9 | 125314173 | 0.135 | 0.019 | 2.83E-12 | 0.111 | 51.9 | 13529 | 1515 | 817 | 17 | -0.010 | 0.010 | 0.30 | 0.003 | 0.47 | -0.078 | 0.074 | 0.29 | 0.46 | -0.096 | 0.024 | 1.04E-04 |
| cg17538538 | 6 | 106825021 | 0.131 | 0.026 | 8.85E-07 | 0.056 | 24.9 | 14420 | 2498 | 1195 | 4 | 0.020 | 0.021 | 0.33 | 0.002 | 0.50 | 0.151 | 0.152 | 0.32 | 0.49 | 0.133 | 0.037 | 3.46E-04 |
| cg23519631 | 12 | 72690355 | 0.111 | 0.022 | 7.26E-07 | 0.057 | 25.3 | 12956 | 1472 | 563 | 8 | -0.016 | 0.016 | 0.33 | 0.002 | 0.50 | -0.144 | 0.144 | 0.32 | 0.49 | -0.164 | 0.034 | 1.82E-06 |
| cg08792976 | 16 | 760528 | 0.103 | 0.011 | 1.48E-19 | 0.179 | 90.6 | 22023 | 2572 | 1696 | 24 | -0.008 | 0.008 | 0.34 | 0.002 | 0.50 | -0.076 | 0.078 | 0.33 | 0.50 | -0.137 | 0.033 | 3.27E-05 |
| cg20623853 | 19 | 1734365 | 0.040 | 0.008 | 2.88E-07 | 0.061 | 27.2 | 17950 | 1888 | 1390 | 13 | -0.006 | 0.006 | 0.34 | 0.002 | 0.50 | -0.132 | 0.135 | 0.33 | 0.50 | -0.148 | 0.036 | 5.67E-05 |
| cg18042910 | 13 | 19662609 | 0.053 | 0.007 | 1.31E-13 | 0.124 | 58.7 | 17061 | 2665 | 1048 | 44 | -0.004 | 0.004 | 0.37 | 0.002 | 0.53 | -0.068 | 0.074 | 0.36 | 0.53 | -0.107 | 0.026 | 4.78E-05 |
| cg14204490 | 8 | 33330967 | 0.035 | 0.006 | 3.81E-10 | 0.090 | 41.2 | 13877 | 1541 | 509 | 27 | 0.006 | 0.007 | 0.37 | 0.002 | 0.53 | 0.176 | 0.204 | 0.39 | 0.55 | -0.222 | 0.056 | 8.23E-05 |
| cg01124546 | 11 | 117515332 | 0.065 | 0.009 | 5.51E-12 | 0.108 | 50.4 | 15525 | 2294 | 1396 | 13 | -0.009 | 0.010 | 0.38 | 0.002 | 0.54 | -0.135 | 0.150 | 0.37 | 0.53 | -0.198 | 0.048 | 4.04E-05 |
| cg00188627 | 3 | 159557779 | 0.133 | 0.022 | 1.50E-09 | 0.084 | 38.2 | 14587 | 1437 | 649 | 8 | 0.012 | 0.014 | 0.40 | 0.002 | 0.56 | 0.089 | 0.104 | 0.39 | 0.55 | 0.105 | 0.030 | 6.05E-04 |
| cg03510150 | 9 | 130690744 | 0.058 | 0.008 | 5.78E-13 | 0.117 | 55.4 | 13396 | 1284 | 730 | 17 | -0.007 | 0.009 | 0.42 | 0.002 | 0.59 | -0.119 | 0.145 | 0.41 | 0.58 | -0.200 | 0.050 | 8.62E-05 |
| cg09701145 | 11 | 2019436 | 0.085 | 0.011 | 7.50E-13 | 0.116 | 54.8 | 17726 | 2869 | | | | | | | | | | | | | | |

SUPPLEMENTARY DATA

| | | | | | | | | | | | | | | | | | | | | | | | |
|------------|----|-----------|-------|-------|------------|-------|-------|-------|------|------|-----|--------|-------|------|-------|------|--------|-------|------|------|--------|-------|----------|
| cg00373987 | 22 | 37868968 | 0.228 | 0.056 | 6.16E-05 | 0.038 | 16.4 | 15697 | 2077 | 1308 | 2 | -0.027 | 0.039 | 0.49 | 0.001 | 0.65 | -0.121 | 0.169 | 0.48 | 0.64 | -0.144 | 0.033 | 1.38E-05 |
| cg12161743 | 11 | 76750531 | 0.055 | 0.007 | 2.42E-14 | 0.131 | 62.5 | 13871 | 1984 | 1045 | 23 | -0.004 | 0.005 | 0.49 | 0.001 | 0.65 | -0.065 | 0.093 | 0.48 | 0.64 | -0.147 | 0.033 | 1.38E-05 |
| cg13827123 | 13 | 93971495 | 0.313 | 0.105 | 0.00304799 | 0.021 | 8.9 | 14374 | 2257 | 949 | 1 | 0.036 | 0.054 | 0.50 | 0.001 | 0.66 | 0.116 | 0.185 | 0.53 | 0.68 | -0.100 | 0.024 | 5.07E-05 |
| cg08491066 | 1 | 156030669 | 0.101 | 0.019 | 1.11E-07 | 0.066 | 29.2 | 11201 | 951 | 464 | 7 | -0.010 | 0.016 | 0.51 | 0.001 | 0.66 | -0.104 | 0.153 | 0.50 | 0.66 | -0.157 | 0.038 | 5.39E-05 |
| cg15575356 | 11 | 47616757 | 0.137 | 0.032 | 1.70E-05 | 0.044 | 18.9 | 11864 | 1306 | 471 | 4 | -0.020 | 0.032 | 0.52 | 0.001 | 0.68 | -0.152 | 0.231 | 0.51 | 0.67 | -0.201 | 0.047 | 2.78E-05 |
| cg00078259 | 6 | 2656232 | 0.148 | 0.020 | 2.76E-13 | 0.121 | 57.0 | 16279 | 2501 | 1372 | 11 | -0.008 | 0.012 | 0.53 | 0.001 | 0.68 | -0.051 | 0.080 | 0.52 | 0.68 | -0.105 | 0.027 | 1.50E-04 |
| cg16140890 | 1 | 1082406 | 0.144 | 0.032 | 1.06E-05 | 0.046 | 19.9 | 16090 | 1384 | 843 | 5 | 0.010 | 0.017 | 0.56 | 0.001 | 0.72 | 0.068 | 0.122 | 0.58 | 0.72 | -0.093 | 0.025 | 2.64E-04 |
| cg21927431 | 9 | 134678366 | 0.173 | 0.028 | 2.39E-09 | 0.082 | 37.2 | 14813 | 1885 | 1114 | 6 | -0.012 | 0.021 | 0.58 | 0.001 | 0.73 | -0.076 | 0.136 | 0.58 | 0.72 | -0.127 | 0.034 | 2.49E-04 |
| cg20659588 | 1 | 237963267 | 0.262 | 0.054 | 1.77E-06 | 0.053 | 23.5 | 15158 | 2388 | 1277 | 3 | -0.018 | 0.034 | 0.59 | 0.001 | 0.73 | -0.073 | 0.132 | 0.58 | 0.72 | -0.125 | 0.029 | 2.48E-05 |
| cg13022891 | 9 | 5817706 | 0.054 | 0.006 | 2.68E-17 | 0.158 | 78.2 | 16090 | 1838 | 1205 | 38 | -0.003 | 0.005 | 0.59 | 0.001 | 0.73 | -0.054 | 0.098 | 0.58 | 0.72 | -0.148 | 0.038 | 1.43E-04 |
| cg25903144 | 7 | 13169086 | 0.086 | 0.017 | 3.43E-07 | 0.061 | 26.9 | 15819 | 2225 | 1281 | 10 | -0.004 | 0.008 | 0.59 | 0.001 | 0.73 | -0.052 | 0.095 | 0.59 | 0.72 | -0.083 | 0.023 | 4.29E-04 |
| cg27410679 | 17 | 43866278 | 0.068 | 0.010 | 3.00E-11 | 0.101 | 46.7 | 12078 | 2955 | 549 | 28 | -0.003 | 0.006 | 0.61 | 0.001 | 0.73 | -0.041 | 0.080 | 0.61 | 0.73 | -0.108 | 0.026 | 5.00E-05 |
| cg10883064 | 17 | 60757346 | 0.085 | 0.014 | 1.18E-09 | 0.085 | 38.8 | 12442 | 1144 | 568 | 20 | -0.004 | 0.009 | 0.62 | 0.001 | 0.73 | -0.050 | 0.098 | 0.61 | 0.73 | -0.110 | 0.030 | 2.51E-04 |
| cg11837181 | 1 | 3001002 | 0.101 | 0.026 | 0.00015292 | 0.034 | 14.6 | 17515 | 2132 | 1321 | 3 | -0.009 | 0.018 | 0.62 | 0.001 | 0.73 | -0.092 | 0.180 | 0.61 | 0.73 | -0.127 | 0.033 | 1.51E-04 |
| cg17985533 | 11 | 2019116 | 0.038 | 0.003 | 8.78E-38 | 0.328 | 202.9 | 17722 | 2867 | 1729 | 108 | -0.001 | 0.003 | 0.62 | 0.001 | 0.73 | -0.038 | 0.076 | 0.62 | 0.73 | -0.175 | 0.043 | 5.97E-05 |
| cg21778193 | 1 | 1098070 | 0.189 | 0.067 | 0.00483001 | 0.019 | 8.0 | 16295 | 1400 | 851 | 1 | -0.018 | 0.040 | 0.66 | 0.000 | 0.77 | -0.087 | 0.192 | 0.65 | 0.76 | -0.119 | 0.029 | 4.25E-05 |
| cg05851077 | 9 | 29802712 | 0.146 | 0.033 | 1.68E-05 | 0.044 | 19.0 | 16112 | 2731 | 1143 | 5 | -0.006 | 0.013 | 0.67 | 0.000 | 0.77 | -0.039 | 0.091 | 0.67 | 0.77 | -0.084 | 0.019 | 1.04E-05 |
| cg06981471 | 7 | 142982076 | 0.125 | 0.026 | 1.98E-06 | 0.053 | 23.3 | 10365 | 1303 | 535 | 5 | 0.008 | 0.020 | 0.67 | 0.000 | 0.77 | 0.064 | 0.155 | 0.68 | 0.77 | -0.127 | 0.036 | 4.27E-04 |
| cg12011250 | 1 | 30198037 | 0.065 | 0.008 | 3.19E-16 | 0.148 | 72.4 | 13795 | 2188 | 1217 | 29 | -0.002 | 0.005 | 0.69 | 0.000 | 0.78 | -0.033 | 0.080 | 0.68 | 0.77 | -0.120 | 0.030 | 6.05E-05 |
| cg07728373 | 2 | 229266790 | 0.077 | 0.016 | 1.82E-06 | 0.053 | 23.4 | 14340 | 2115 | 872 | 11 | -0.003 | 0.007 | 0.69 | 0.000 | 0.78 | -0.039 | 0.096 | 0.68 | 0.77 | -0.108 | 0.021 | 5.53E-07 |
| cg16608656 | 11 | 1082797 | 0.103 | 0.033 | 0.00173057 | 0.023 | 9.9 | 16848 | 2415 | 1343 | 2 | -0.008 | 0.020 | 0.70 | 0.000 | 0.78 | -0.072 | 0.184 | 0.69 | 0.78 | -0.112 | 0.029 | 1.47E-04 |
| cg13625403 | 11 | 120088510 | 0.057 | 0.012 | 3.76E-06 | 0.050 | 22.0 | 15102 | 2093 | 1233 | 9 | 0.003 | 0.007 | 0.71 | 0.000 | 0.79 | 0.047 | 0.129 | 0.71 | 0.79 | -0.117 | 0.028 | 4.48E-05 |
| cg00918900 | 1 | 1057335 | 0.087 | 0.012 | 1.22E-12 | 0.114 | 53.7 | 15892 | 1367 | 827 | 21 | -0.003 | 0.007 | 0.71 | 0.000 | 0.79 | -0.031 | 0.083 | 0.71 | 0.79 | -0.124 | 0.029 | 1.83E-05 |
| cg00403246 | 10 | 134794753 | 0.096 | 0.023 | 3.36E-05 | 0.041 | 17.6 | 18305 | 1964 | 1252 | 2 | 0.009 | 0.025 | 0.73 | 0.000 | 0.79 | 0.090 | 0.263 | 0.73 | 0.80 | -0.188 | 0.051 | 2.46E-04 |
| cg11138231 | 7 | 99475824 | 0.111 | 0.017 | 4.91E-11 | 0.099 | 45.6 | 12236 | 880 | 344 | 20 | -0.003 | 0.008 | 0.76 | 0.000 | 0.83 | -0.022 | 0.073 | 0.76 | 0.82 | -0.095 | 0.023 | 4.48E-05 |
| cg08474716 | 16 | 27079640 | 0.206 | 0.029 | 4.42E-12 | 0.109 | 50.9 | 16195 | 2472 | 1481 | 8 | -0.004 | 0.016 | 0.78 | 0.000 | 0.84 | -0.022 | 0.078 | 0.78 | 0.83 | -0.092 | 0.025 | 2.08E-04 |
| cg07203271 | 1 | 53926112 | 0.145 | 0.024 | 2.03E-09 | 0.083 | 37.6 | 14665 | 2120 | 967 | 9 | -0.004 | 0.015 | 0.78 | 0.000 | 0.84 | -0.028 | 0.101 | 0.78 | 0.83 | -0.104 | 0.029 | 4.27E-04 |
| cg25380707 | 15 | 86660946 | 0.252 | 0.080 | 0.00180737 | 0.023 | 9.9 | 17604 | 2562 | 1510 | 1 | -0.010 | 0.040 | 0.81 | 0.000 | 0.85 | -0.043 | 0.176 | 0.81 | 0.85 | -0.086 | 0.024 | 2.83E-04 |
| cg17905524 | 15 | 28305412 | 0.106 | 0.014 | 1.08E-12 | 0.115 | 54.0 | 11317 | 1447 | 661 | 17 | -0.002 | 0.010 | 0.82 | 0.000 | 0.86 | -0.021 | 0.094 | 0.82 | 0.86 | -0.127 | 0.030 | 2.77E-05 |
| cg18616558 | 7 | 158798592 | 0.080 | 0.014 | 1.49E-08 | 0.074 | 33.4 | 14247 | 1918 | 902 | 20 | -0.001 | 0.006 | 0.92 | 0.000 | 0.95 | -0.008 | 0.073 | 0.92 | 0.95 | -0.071 | 0.020 | 4.52E-04 |
| cg04140215 | 2 | 60540175 | 0.210 | 0.075 | 0.00563409 | 0.018 | 7.7 | 15077 | 1764 | 1102 | 1 | 0.004 | 0.050 | 0.94 | 0.000 | 0.96 | 0.021 | 0.265 | 0.94 | 0.97 | -0.119 | 0.031 | 1.48E-04 |
| cg26238771 | 19 | 592158 | 0.117 | 0.011 | 3.43E-23 | 0.211 | 111.2 | 17193 | 2642 | 2036 | 20 | 0.000 | 0.009 | 0.98 | 0.000 | 0.99 | -0.002 | 0.080 | 0.98 | 0.99 | -0.139 | 0.036 | 1.21E-04 |
| cg05347757 | 15 | 51121020 | 0.103 | 0.034 | 0.00251697 | 0.022 | 9.2 | 13001 | 1995 | 637 | 3 | 0.000 | 0.018 | 0.98 | 0.000 | 0.99 | -0.004 | 0.173 | 0.98 | 0.99 | -0.083 | 0.025 | 1.10E-03 |
| cg02688867 | 12 | 133019854 | 0.180 | 0.055 | 0.0012479 | 0.025 | 10.6 | 16623 | 2355 | 1394 | 2 | -0.001 | 0.033 | 0.98 | 0.000 | 0.99 | -0.004 | 0.179 | 0.98 | 0.99 | -0.100 | 0.028 | 4.92E-04 |
| cg15952840 | 8 | 11451312 | 0.159 | 0.035 | 5.66E-06 | 0.048 | 21.1 | 21931 | 3018 | 2084 | 3 | 0.000 | 0.022 | 0.99 | 0.000 | 0.99 | 0.003 | 0.145 | 0.99 | 0.99 | -0.128 | 0.030 | 2.32E-05 |
| cg00045690 | 6 | 32825897 | | | | | | 19525 | 8434 | 2923 | 0 | | | | | | | | | | -0.086 | 0.022 | 1.12E-04 |
| cg01216882 | 20 | 31804277 | | | | | | 14128 | 1327 | 702 | 0 | | | | | | | | | | -0.137 | 0.031 | 9.94E-06 |
| cg01530082 | 6 | 32784327 | | | | | | 19703 | 8668 | 2934 | 0 | | | | | | | | | | -0.107 | 0.026 | 4.05E-05 |
| cg01615861 | 16 | 85513978 | | | | | | 21138 | 2881 | 2231 | 0 | | | | | | | | | | -0.283 | 0.080 | 4.45E-04 |
| cg02831474 | 10 | 1783879 | | | | | | 18860 | 3144 | 1828 | 0 | | | | | | | | | | -0.113 | 0.024 | 3.12E-06 |
| cg03275961 | 9 | 90386329 | | | | | | 15207 | 2428 | 1223 | 0 | | | | | | | | | | -0.161 | 0.040 | 5.57E-05 |
| cg03710412 | 8 | 142938779 | | | | | | 17500 | 2608 | 1446 | 0 | | | | | | | | | | -0.091 | 0.026 | 4.85E-04 |
| cg04847540 | 1 | 182754070 | | | | | | 13816 | 1976 | 812 | 0 | | | | | | | | | | -0.112 | 0.028 | 7.64E-05 |
| cg04907218 | 12 | 132826672 | | | | | | 16467 | 2098 | 1300 | 0 | | | | | | | | | | -0.118 | 0.030 | 9.09E-05 |
| cg05444304 | 2 | 31442675 | | | | | | 15973 | 2123 | 1458 | 0 | | | | | | | | | | 0.133 | 0.039 | 6.39E-04 |
| cg05790369 | 17 | 33202007 | | | | | | 14788 | 2007 | 1086 | 0 | | | | | | | | | | -0.115 | 0.030 | 1.90E-04 |
| cg06343669 | 12 | 132690782 | | | | | | 16014 | 2044 | 1320 | 0 | | | | | | | | | | -0.117 | 0.028 | 2.72E-05 |
| cg06495184 | 19 | 406256 | | | | | | 13720 | 2023 | 1579 | 0 | | | | | | | | | | -0.142 | 0.035 | 5.34E-05 |
| cg06830529 | 9 | 124960107 | | | | | | 14549 | 1938 | 1035 | 0 | | | | | | | | | | -0.139 | 0.037 | 1.73E-04 |
| cg07080246 | 13 | 63306618 | | | | | | 13919 | 2164 | 705 | 0 | | | | | | | | | | -0.076 | 0.020 | 1.22E-04 |
| cg07686906 | 1 | 3353825 | | | | | | 18958 | 2698 | 1695 | 0 | | | | | | | | | | -0.150 | 0.037 | 5.70E-05 |
| cg07865738 | 2 | 3649447 | | | | | | 15986 | 2592 | 1436 | 0 | | | | | | | | | | -0.095 | 0.024 | 6.57E-05 |

SUPPLEMENTARY DATA

| | | | | | | | | | | | | | | | | | | | | |
|------------|----|-----------|--|--|--|--|--|-------|------|------|---|--|--|--|--|--|--|--------|-------|----------|
| cg08058801 | 3 | 147974120 | | | | | | 14142 | 1723 | 693 | 0 | | | | | | | -0.114 | 0.026 | 1.13E-05 |
| cg08357744 | 17 | 37559321 | | | | | | 13259 | 1346 | 609 | 0 | | | | | | | -0.084 | 0.023 | 2.31E-04 |
| cg08372350 | 16 | 88943191 | | | | | | 22238 | 2702 | 1867 | 0 | | | | | | | -0.107 | 0.028 | 1.59E-04 |
| cg08389995 | 3 | 131645295 | | | | | | 14602 | 2835 | 1125 | 0 | | | | | | | -0.083 | 0.023 | 3.49E-04 |
| cg09181711 | 19 | 51327196 | | | | | | 16264 | 2335 | 1602 | 0 | | | | | | | -0.136 | 0.029 | 2.61E-06 |
| cg09851596 | 8 | 143545214 | | | | | | 18124 | 2792 | 1362 | 0 | | | | | | | -0.177 | 0.044 | 7.97E-05 |
| cg10877514 | 17 | 292539 | | | | | | 12641 | 1872 | 1292 | 0 | | | | | | | -0.105 | 0.028 | 1.79E-04 |
| cg11086223 | 18 | 11487579 | | | | | | 16274 | 2395 | 1276 | 0 | | | | | | | -0.135 | 0.034 | 6.79E-05 |
| cg11177743 | 20 | 21545361 | | | | | | 13270 | 1160 | 444 | 0 | | | | | | | -0.138 | 0.033 | 3.15E-05 |
| cg11294782 | 3 | 10487409 | | | | | | 15223 | 2264 | 1314 | 0 | | | | | | | -0.152 | 0.040 | 1.69E-04 |
| cg11416433 | 15 | 25429261 | | | | | | 16557 | 1988 | 1141 | 0 | | | | | | | -0.201 | 0.044 | 6.72E-06 |
| cg11821795 | 22 | 47611604 | | | | | | 19073 | 2440 | 1408 | 0 | | | | | | | -0.109 | 0.027 | 4.59E-05 |
| cg12697796 | 10 | 49954016 | | | | | | 15686 | 2331 | 1086 | 0 | | | | | | | -0.179 | 0.041 | 2.03E-05 |
| cg13116906 | 12 | 119552989 | | | | | | 14166 | 2319 | 1181 | 0 | | | | | | | -0.105 | 0.027 | 1.44E-04 |
| cg13254480 | 16 | 86513248 | | | | | | 23316 | 3570 | 2722 | 0 | | | | | | | -0.073 | 0.020 | 2.91E-04 |
| cg13257690 | 14 | 101908752 | | | | | | 15386 | 2170 | 1265 | 0 | | | | | | | -0.178 | 0.046 | 1.17E-04 |
| cg13480617 | 5 | 35230725 | | | | | | 14639 | 2107 | 978 | 0 | | | | | | | -0.121 | 0.038 | 1.49E-03 |
| cg14494450 | 1 | 4190881 | | | | | | 15856 | 2958 | 1567 | 0 | | | | | | | -0.116 | 0.027 | 2.07E-05 |
| cg14757962 | 5 | 2176287 | | | | | | 19312 | 3591 | 2168 | 0 | | | | | | | -0.108 | 0.030 | 3.06E-04 |
| cg14759277 | 17 | 48674158 | | | | | | 13790 | 1755 | 1128 | 0 | | | | | | | -0.164 | 0.041 | 7.62E-05 |
| cg14886500 | 6 | 23447204 | | | | | | 17748 | 2646 | 1490 | 0 | | | | | | | -0.080 | 0.023 | 7.30E-04 |
| cg16513326 | 8 | 10877312 | | | | | | 21573 | 3120 | 2033 | 0 | | | | | | | -0.111 | 0.032 | 5.03E-04 |
| cg16857113 | 3 | 1133865 | | | | | | 18339 | 2963 | 1779 | 0 | | | | | | | -0.114 | 0.029 | 1.12E-04 |
| cg17224410 | 9 | 137852661 | | | | | | 19804 | 2883 | 1965 | 0 | | | | | | | -0.123 | 0.033 | 2.00E-04 |
| cg17226458 | 6 | 33780022 | | | | | | 16101 | 2493 | 1395 | 0 | | | | | | | -0.145 | 0.039 | 2.71E-04 |
| cg17587481 | 5 | 152603672 | | | | | | 14222 | 2224 | 764 | 0 | | | | | | | -0.103 | 0.024 | 1.78E-05 |
| cg18108190 | 19 | 2727822 | | | | | | 18307 | 2844 | 1847 | 0 | | | | | | | -0.142 | 0.033 | 2.05E-05 |
| cg18233010 | 10 | 114911413 | | | | | | 14248 | 1705 | 1135 | 0 | | | | | | | 0.170 | 0.045 | 2.13E-04 |
| cg18617770 | 12 | 133035939 | | | | | | 16438 | 2356 | 1404 | 0 | | | | | | | -0.160 | 0.037 | 2.04E-05 |
| cg18650968 | 4 | 176340837 | | | | | | 14564 | 1524 | 726 | 0 | | | | | | | -0.100 | 0.028 | 4.36E-04 |
| cg19450046 | 21 | 45712938 | | | | | | 18589 | 2432 | 1382 | 0 | | | | | | | -0.138 | 0.036 | 1.66E-04 |
| cg20112984 | 10 | 1447183 | | | | | | 18134 | 2767 | 1723 | 0 | | | | | | | -0.103 | 0.027 | 2.03E-04 |
| cg21307723 | 1 | 245215135 | | | | | | 15619 | 2326 | 1412 | 0 | | | | | | | 0.143 | 0.038 | 2.22E-04 |
| cg23016281 | 22 | 27616620 | | | | | | 17311 | 3036 | 1986 | 0 | | | | | | | -0.092 | 0.025 | 2.60E-04 |
| cg23359229 | 13 | 29976597 | | | | | | 15238 | 2418 | 1020 | 0 | | | | | | | -0.110 | 0.026 | 2.82E-05 |
| cg23536293 | 12 | 99524459 | | | | | | 13533 | 1605 | 925 | 0 | | | | | | | -0.071 | 0.021 | 6.41E-04 |
| cg23561197 | 5 | 125852064 | | | | | | 15210 | 2226 | 1108 | 0 | | | | | | | -0.106 | 0.030 | 4.06E-04 |
| cg25314607 | 5 | 2148308 | | | | | | 19252 | 3620 | 2202 | 0 | | | | | | | -0.122 | 0.030 | 6.85E-05 |
| cg26178126 | 1 | 37740931 | | | | | | 14242 | 2023 | 983 | 0 | | | | | | | -0.125 | 0.031 | 6.10E-05 |
| cg27332583 | 18 | 35150602 | | | | | | 13359 | 1749 | 916 | 0 | | | | | | | -0.175 | 0.041 | 2.12E-05 |

SUPPLEMENTARY DATA

Supplementary Table 5. Results from Mendelian randomization analyses using GRS representing insulin sensitivity (8 SNPs) associations with methylation levels at CpG sites identified in Model 1

| CpG | chr | pos | Maternal GRS8 (raw) association with placenta methylation | | | |
|------------|-------|-----------|---|-------|---------|---------|
| | | | estimate | SE | P_value | Rsquare |
| cg12161743 | chr11 | 76750531 | 0.048 | 0.022 | 0.03 | 0.0112 |
| cg13625403 | chr11 | 120008510 | -0.041 | 0.026 | 0.12 | 0.0059 |
| cg04293688 | chr2 | 75466188 | -0.039 | 0.026 | 0.13 | 0.0056 |
| cg01216882 | chr20 | 31804277 | 0.035 | 0.024 | 0.15 | 0.0051 |
| cg13254480 | chr16 | 86513248 | 0.053 | 0.038 | 0.16 | 0.0047 |
| cg26225794 | chr7 | 27717723 | -0.030 | 0.023 | 0.19 | 0.0042 |
| cg13022891 | chr9 | 5817706 | -0.024 | 0.019 | 0.22 | 0.0036 |
| cg06830529 | chr9 | 124960107 | 0.025 | 0.021 | 0.23 | 0.0034 |
| cg26178126 | chr1 | 37740931 | 0.029 | 0.024 | 0.24 | 0.0034 |
| cg09063835 | chr20 | 58181392 | 0.032 | 0.027 | 0.25 | 0.0032 |
| cg23519631 | chr12 | 72690355 | 0.026 | 0.022 | 0.25 | 0.0032 |
| cg16853336 | chr3 | 149396490 | -0.016 | 0.015 | 0.29 | 0.0026 |
| cg07080246 | chr13 | 63306618 | -0.038 | 0.038 | 0.32 | 0.0023 |
| cg12760575 | chr9 | 139158654 | -0.019 | 0.019 | 0.33 | 0.0023 |
| cg07435932 | chr13 | 19648633 | 0.030 | 0.031 | 0.33 | 0.0023 |
| cg06343669 | chr12 | 132690782 | -0.025 | 0.027 | 0.36 | 0.0020 |
| cg24848724 | chr8 | 142578640 | 0.028 | 0.031 | 0.36 | 0.0020 |
| cg18616558 | chr7 | 158798592 | 0.033 | 0.037 | 0.37 | 0.0019 |
| cg18617770 | chr12 | 133035939 | -0.018 | 0.020 | 0.38 | 0.0019 |
| cg07728373 | chr2 | 229266790 | -0.031 | 0.035 | 0.38 | 0.0018 |
| cg03275961 | chr9 | 90386329 | 0.016 | 0.019 | 0.40 | 0.0017 |
| cg16340422 | chr17 | 17110120 | 0.021 | 0.026 | 0.41 | 0.0016 |
| cg00903007 | chr17 | 34068171 | -0.011 | 0.014 | 0.42 | 0.0016 |
| cg06077821 | chr5 | 180048893 | -0.017 | 0.022 | 0.43 | 0.0015 |
| cg12011250 | chr1 | 30198037 | -0.020 | 0.025 | 0.43 | 0.0015 |
| cg09640571 | chr16 | 88388040 | 0.012 | 0.015 | 0.43 | 0.0015 |
| cg11821795 | chr22 | 47611604 | 0.021 | 0.028 | 0.46 | 0.0013 |
| cg09181711 | chr19 | 51327196 | -0.018 | 0.026 | 0.48 | 0.0012 |
| cg24738497 | chr19 | 36005395 | -0.016 | 0.022 | 0.48 | 0.0012 |
| cg17587481 | chr5 | 152603672 | -0.021 | 0.031 | 0.50 | 0.0011 |
| cg00460471 | chr2 | 61437605 | -0.018 | 0.027 | 0.50 | 0.0011 |
| cg18216439 | chr18 | 35132882 | 0.017 | 0.025 | 0.50 | 0.0011 |
| cg14440140 | chr9 | 140177567 | 0.017 | 0.025 | 0.51 | 0.0010 |
| cg12697796 | chr10 | 49954016 | 0.012 | 0.018 | 0.51 | 0.0010 |
| cg00386725 | chr1 | 18617791 | 0.011 | 0.017 | 0.52 | 0.0010 |
| cg05790369 | chr17 | 33202007 | 0.016 | 0.025 | 0.52 | 0.0010 |
| cg27410679 | chr17 | 43866278 | 0.018 | 0.029 | 0.53 | 0.0010 |
| cg08792976 | chr16 | 760528 | -0.014 | 0.023 | 0.53 | 0.0009 |
| cg10877514 | chr17 | 292539 | 0.017 | 0.027 | 0.53 | 0.0009 |
| cg18450810 | chr17 | 51111066 | 0.016 | 0.026 | 0.53 | 0.0009 |
| cg06066032 | chr13 | 48369175 | 0.019 | 0.031 | 0.54 | 0.0009 |
| cg18861231 | chr3 | 152057943 | -0.018 | 0.030 | 0.54 | 0.0009 |
| cg10964639 | chr16 | 24043296 | 0.014 | 0.024 | 0.55 | 0.0008 |
| cg02384870 | chr2 | 216161208 | 0.018 | 0.031 | 0.56 | 0.0008 |
| cg17226458 | chr6 | 33780022 | 0.011 | 0.019 | 0.56 | 0.0008 |
| cg00373987 | chr22 | 37868968 | -0.013 | 0.023 | 0.57 | 0.0008 |
| cg01864178 | chr20 | 44662890 | -0.014 | 0.026 | 0.58 | 0.0008 |
| cg24073146 | chr11 | 2543204 | -0.010 | 0.018 | 0.58 | 0.0007 |
| cg24475484 | chr8 | 1625395 | 0.016 | 0.030 | 0.58 | 0.0007 |
| cg27652925 | chr6 | 31740951 | 0.008 | 0.014 | 0.59 | 0.0007 |
| cg23536293 | chr12 | 99524459 | 0.020 | 0.037 | 0.60 | 0.0007 |
| cg05808230 | chr20 | 61726456 | 0.012 | 0.023 | 0.60 | 0.0007 |
| cg21396983 | chr16 | 1275423 | 0.013 | 0.025 | 0.61 | 0.0006 |
| cg25380707 | chr15 | 86660946 | 0.016 | 0.032 | 0.62 | 0.0006 |
| cg09701145 | chr11 | 2019436 | -0.010 | 0.019 | 0.62 | 0.0006 |
| cg08058801 | chr3 | 147974120 | 0.015 | 0.029 | 0.62 | 0.0006 |
| cg15952840 | chr8 | 11451312 | 0.012 | 0.025 | 0.62 | 0.0006 |
| cg08582801 | chr7 | 99588335 | 0.007 | 0.013 | 0.62 | 0.0006 |
| cg00918900 | chr1 | 1057335 | 0.012 | 0.026 | 0.63 | 0.0005 |
| cg12658750 | chr18 | 75499382 | 0.009 | 0.019 | 0.63 | 0.0005 |
| cg15575356 | chr11 | 47616757 | -0.008 | 0.016 | 0.64 | 0.0005 |
| cg20659588 | chr1 | 237963267 | -0.012 | 0.025 | 0.64 | 0.0005 |
| cg02287371 | chr5 | 1246274 | -0.013 | 0.027 | 0.64 | 0.0005 |
| cg16140890 | chr1 | 1082406 | 0.014 | 0.030 | 0.65 | 0.0005 |
| cg26849382 | chr5 | 140901022 | 0.013 | 0.030 | 0.65 | 0.0005 |
| cg03710412 | chr8 | 142938779 | -0.013 | 0.029 | 0.66 | 0.0005 |
| cg20623853 | chr19 | 1734365 | 0.009 | 0.021 | 0.66 | 0.0005 |
| cg01124546 | chr11 | 117515332 | -0.007 | 0.016 | 0.66 | 0.0005 |
| cg26238771 | chr19 | 592158 | 0.009 | 0.021 | 0.66 | 0.0005 |
| cg08389995 | chr3 | 131645295 | -0.014 | 0.033 | 0.66 | 0.0005 |
| cg13257690 | chr14 | 101908752 | 0.007 | 0.016 | 0.67 | 0.0004 |
| cg17756860 | chr4 | 110510374 | -0.011 | 0.027 | 0.67 | 0.0004 |

SUPPLEMENTARY DATA

| | | | | | | |
|------------|-------|-----------|--------|-------|------|--------|
| cg01530082 | chr6 | 32784327 | -0.012 | 0.029 | 0.67 | 0.0004 |
| cg00045690 | chr6 | 32825897 | 0.014 | 0.034 | 0.67 | 0.0004 |
| cg14839558 | chr1 | 1874488 | 0.012 | 0.028 | 0.67 | 0.0004 |
| cg25314607 | chr5 | 2148308 | 0.011 | 0.025 | 0.67 | 0.0004 |
| cg00781180 | chr16 | 73984002 | -0.011 | 0.026 | 0.67 | 0.0004 |
| cg08474716 | chr16 | 27079640 | 0.013 | 0.031 | 0.67 | 0.0004 |
| cg13480617 | chr5 | 35230725 | 0.008 | 0.020 | 0.67 | 0.0004 |
| cg11416433 | chr15 | 25429261 | 0.007 | 0.017 | 0.67 | 0.0004 |
| cg17538538 | chr6 | 106825021 | 0.009 | 0.020 | 0.67 | 0.0004 |
| cg05775586 | chr9 | 28889354 | -0.012 | 0.029 | 0.68 | 0.0004 |
| cg08737919 | chr12 | 108840669 | 0.010 | 0.026 | 0.69 | 0.0004 |
| cg01618245 | chr20 | 61990279 | 0.012 | 0.030 | 0.69 | 0.0004 |
| cg21307723 | chr1 | 245215135 | 0.008 | 0.019 | 0.69 | 0.0004 |
| cg20018106 | chr4 | 21768009 | -0.011 | 0.028 | 0.70 | 0.0004 |
| cg02831474 | chr10 | 1783879 | 0.012 | 0.031 | 0.70 | 0.0004 |
| cg04535454 | chr8 | 144035192 | 0.008 | 0.022 | 0.70 | 0.0004 |
| cg08491066 | chr1 | 156030669 | 0.007 | 0.019 | 0.70 | 0.0003 |
| cg12673377 | chr7 | 1388271 | 0.008 | 0.021 | 0.70 | 0.0003 |
| cg07203271 | chr1 | 53926112 | -0.010 | 0.026 | 0.71 | 0.0003 |
| cg09319487 | chr20 | 56771178 | 0.009 | 0.025 | 0.71 | 0.0003 |
| cg23359229 | chr13 | 29976597 | 0.011 | 0.029 | 0.71 | 0.0003 |
| cg23521973 | chr10 | 125117429 | 0.009 | 0.024 | 0.72 | 0.0003 |
| cg01948226 | chr6 | 32829208 | 0.009 | 0.025 | 0.72 | 0.0003 |
| cg18184975 | chr20 | 3644192 | 0.008 | 0.023 | 0.72 | 0.0003 |
| cg13116906 | chr12 | 119552989 | 0.010 | 0.027 | 0.73 | 0.0003 |
| cg04131502 | chr1 | 2845583 | 0.008 | 0.023 | 0.73 | 0.0003 |
| cg04907218 | chr12 | 132826672 | -0.009 | 0.025 | 0.74 | 0.0003 |
| cg21778193 | chr1 | 1098070 | 0.009 | 0.026 | 0.74 | 0.0003 |
| cg11855710 | chr19 | 29768180 | -0.008 | 0.024 | 0.75 | 0.0002 |
| cg26037026 | chr7 | 1641922 | -0.006 | 0.020 | 0.75 | 0.0002 |
| cg25903144 | chr7 | 131609806 | 0.010 | 0.033 | 0.76 | 0.0002 |
| cg13377560 | chr22 | 43860820 | -0.006 | 0.021 | 0.76 | 0.0002 |
| cg16444062 | chr16 | 2770298 | 0.009 | 0.029 | 0.76 | 0.0002 |
| cg27625886 | chr20 | 62060443 | -0.008 | 0.029 | 0.77 | 0.0002 |
| cg01898225 | chr15 | 99975141 | -0.006 | 0.019 | 0.77 | 0.0002 |
| cg07865738 | chr2 | 3649447 | -0.009 | 0.032 | 0.78 | 0.0002 |
| cg23561197 | chr5 | 125852064 | -0.007 | 0.025 | 0.79 | 0.0002 |
| cg14361947 | chr9 | 140780551 | 0.004 | 0.015 | 0.79 | 0.0002 |
| cg08099672 | chr9 | 139944162 | -0.005 | 0.021 | 0.79 | 0.0002 |
| cg08357744 | chr17 | 37559321 | 0.008 | 0.034 | 0.80 | 0.0002 |
| cg03568769 | chr11 | 1800837 | 0.006 | 0.023 | 0.80 | 0.0001 |
| cg16857113 | chr3 | 1133865 | -0.006 | 0.026 | 0.81 | 0.0001 |
| cg18682781 | chr5 | 176277687 | 0.007 | 0.027 | 0.81 | 0.0001 |
| cg25721832 | chr3 | 81913557 | -0.006 | 0.023 | 0.81 | 0.0001 |
| cg27332583 | chr18 | 35150602 | -0.004 | 0.019 | 0.81 | 0.0001 |
| cg11294782 | chr3 | 10487409 | 0.004 | 0.019 | 0.82 | 0.0001 |
| cg08511578 | chr10 | 1403938 | -0.007 | 0.030 | 0.82 | 0.0001 |
| cg07686906 | chr1 | 3353825 | 0.005 | 0.020 | 0.82 | 0.0001 |
| cg23016281 | chr22 | 27616620 | -0.007 | 0.030 | 0.83 | 0.0001 |
| cg08149223 | chr10 | 50507130 | -0.004 | 0.016 | 0.83 | 0.0001 |
| cg24600306 | chr20 | 61990671 | 0.004 | 0.016 | 0.83 | 0.0001 |
| cg02688867 | chr12 | 133019854 | -0.006 | 0.026 | 0.83 | 0.0001 |
| cg05444304 | chr2 | 31442675 | 0.004 | 0.020 | 0.83 | 0.0001 |
| cg05389237 | chr9 | 125314173 | 0.006 | 0.031 | 0.84 | 0.0001 |
| cg17985533 | chr11 | 2019116 | 0.003 | 0.017 | 0.84 | 0.0001 |
| cg00389386 | chr1 | 6188136 | 0.003 | 0.017 | 0.84 | 0.0001 |
| cg13827123 | chr13 | 93971495 | -0.006 | 0.031 | 0.85 | 0.0001 |
| cg17905524 | chr15 | 28305412 | 0.005 | 0.025 | 0.85 | 0.0001 |
| cg11177743 | chr20 | 21545361 | 0.004 | 0.023 | 0.86 | 0.0001 |
| cg00046877 | chr22 | 19702664 | 0.002 | 0.013 | 0.86 | 0.0001 |
| cg11138231 | chr7 | 99475824 | 0.006 | 0.033 | 0.86 | 0.0001 |
| cg06981471 | chr7 | 142982076 | -0.004 | 0.021 | 0.86 | 0.0001 |
| cg14757962 | chr5 | 2176287 | -0.004 | 0.025 | 0.87 | 0.0001 |
| cg00078259 | chr6 | 2656232 | 0.004 | 0.027 | 0.87 | 0.0001 |
| cg00188627 | chr3 | 15955779 | 0.004 | 0.025 | 0.88 | 0.0001 |
| cg08372350 | chr16 | 88943191 | -0.004 | 0.027 | 0.88 | 0.0001 |
| cg14886500 | chr6 | 23447204 | -0.005 | 0.032 | 0.88 | 0.0001 |
| cg06751809 | chr3 | 105047227 | 0.005 | 0.037 | 0.88 | 0.0001 |
| cg17224410 | chr9 | 137852661 | -0.003 | 0.023 | 0.89 | 0.0000 |
| cg13449257 | chr9 | 137355128 | 0.003 | 0.022 | 0.89 | 0.0000 |
| cg18650968 | chr4 | 176340837 | -0.004 | 0.027 | 0.89 | 0.0000 |
| cg06429120 | chr16 | 86309214 | 0.005 | 0.036 | 0.89 | 0.0000 |
| cg24857040 | chr13 | 113565214 | -0.003 | 0.024 | 0.89 | 0.0000 |
| cg16513326 | chr8 | 10877312 | -0.003 | 0.024 | 0.89 | 0.0000 |
| cg20112984 | chr10 | 1447183 | -0.004 | 0.028 | 0.90 | 0.0000 |
| cg16608656 | chr11 | 1082797 | -0.003 | 0.026 | 0.90 | 0.0000 |

SUPPLEMENTARY DATA

| | | | | | | |
|------------|-------|-----------|--------|-------|------|--------|
| cg19450046 | chr21 | 45712938 | -0.003 | 0.021 | 0.90 | 0.0000 |
| cg09073467 | chr1 | 6155515 | -0.002 | 0.016 | 0.91 | 0.0000 |
| cg08204714 | chr16 | 87726461 | -0.002 | 0.015 | 0.91 | 0.0000 |
| cg14494450 | chr1 | 4190881 | -0.003 | 0.028 | 0.92 | 0.0000 |
| cg00403246 | chr10 | 134794753 | 0.002 | 0.015 | 0.92 | 0.0000 |
| cg21927431 | chr9 | 134678366 | -0.002 | 0.022 | 0.92 | 0.0000 |
| cg03510150 | chr9 | 130690744 | -0.002 | 0.015 | 0.92 | 0.0000 |
| cg14204490 | chr8 | 33330967 | 0.001 | 0.013 | 0.92 | 0.0000 |
| cg11086223 | chr18 | 11487579 | 0.002 | 0.022 | 0.93 | 0.0000 |
| cg21347874 | chr17 | 77122733 | 0.002 | 0.026 | 0.93 | 0.0000 |
| cg20612875 | chr17 | 1094308 | 0.001 | 0.015 | 0.94 | 0.0000 |
| cg14759277 | chr17 | 48674158 | -0.001 | 0.018 | 0.94 | 0.0000 |
| cg05851077 | chr9 | 29802712 | 0.003 | 0.040 | 0.94 | 0.0000 |
| cg05202700 | chr8 | 38258250 | -0.002 | 0.028 | 0.94 | 0.0000 |
| cg11837181 | chr1 | 3001002 | -0.002 | 0.023 | 0.94 | 0.0000 |
| cg04847540 | chr1 | 182754070 | -0.002 | 0.027 | 0.95 | 0.0000 |
| cg08239663 | chr16 | 52688569 | -0.002 | 0.032 | 0.95 | 0.0000 |
| cg13463516 | chr9 | 138632121 | -0.001 | 0.021 | 0.95 | 0.0000 |
| cg22509138 | chr20 | 44809026 | -0.002 | 0.029 | 0.95 | 0.0000 |
| cg04140215 | chr2 | 60540175 | 0.001 | 0.024 | 0.96 | 0.0000 |
| cg10883064 | chr17 | 60757346 | -0.001 | 0.025 | 0.96 | 0.0000 |
| cg15867652 | chr17 | 79011277 | -0.002 | 0.029 | 0.96 | 0.0000 |
| cg01095004 | chr16 | 88454883 | 0.001 | 0.027 | 0.96 | 0.0000 |
| cg09851596 | chr8 | 143545214 | 0.001 | 0.017 | 0.97 | 0.0000 |
| cg24064348 | chr1 | 54181174 | 0.000 | 0.013 | 0.97 | 0.0000 |
| cg18042910 | chr13 | 19662609 | 0.001 | 0.029 | 0.98 | 0.0000 |
| cg01615861 | chr16 | 85513978 | 0.000 | 0.009 | 0.98 | 0.0000 |
| cg00875983 | chr11 | 3240213 | 0.000 | 0.016 | 0.98 | 0.0000 |
| cg06495184 | chr19 | 406256 | -0.001 | 0.021 | 0.98 | 0.0000 |
| cg18233010 | chr10 | 114911413 | 0.000 | 0.017 | 0.98 | 0.0000 |
| cg12193237 | chr19 | 3907913 | 0.000 | 0.021 | 0.98 | 0.0000 |
| cg16819051 | chr2 | 229173235 | 0.000 | 0.025 | 0.99 | 0.0000 |
| cg11765907 | chr20 | 43103949 | 0.000 | 0.015 | 0.99 | 0.0000 |
| cg25285919 | chr3 | 188312215 | 0.000 | 0.024 | 0.99 | 0.0000 |
| cg11586089 | chr14 | 66644923 | 0.000 | 0.025 | 0.99 | 0.0000 |
| cg03699074 | chr16 | 88849875 | 0.000 | 0.030 | 0.99 | 0.0000 |
| cg18108190 | chr19 | 2727822 | 0.000 | 0.023 | 0.99 | 0.0000 |
| cg05347757 | chr15 | 51121020 | 0.000 | 0.030 | 1.00 | 0.0000 |
| cg12954267 | chr6 | 33093109 | 0.000 | 0.030 | 1.00 | 0.0000 |
| cg02110836 | chr17 | 71953083 | 0.000 | 0.013 | 1.00 | 0.0000 |

SUPPLEMENTARY DATA

Supplementary Table 6: Results from enrichment in TargetScan microRNA 2017 database (178 unique genes from annotation to CpG identified in Model 1)

| Term | Overlap | P-value | Adjusted P-value | Old P-value | Old Adjusted P-value | Z-score | Combined Score | Genes |
|-----------------|---------|-----------|------------------|-------------|----------------------|---------|----------------|---|
| hsa-miR-1538 | 28/1027 | 1.194E-07 | 4.05E-05 | 1.236E-09 | 4.1885E-07 | -1.924 | 30.67 | 16;CTSD;GPC6;TBC1D22A;SLC39A3;NKX2-2;SRRM4;SLC12A5;RBFOX3;PRKCB;RASL10B;IRX4;MTUS2;KCNAB2;BAIAP2;C1ORF159; |
| hsa-miR-4745-3p | 28/1027 | 1.194E-07 | 4.05E-05 | 1.236E-09 | 4.1885E-07 | -1.917 | 30.56 | 16;CTSD;GPC6;TBC1D22A;SLC39A3;NKX2-2;SRRM4;SLC12A5;RBFOX3;PRKCB;RASL10B;IRX4;MTUS2;KCNAB2;BAIAP2;C1ORF159; |
| hsa-miR-3180 | 32/1475 | 2.206E-06 | 0.000299 | 1.538E-08 | 2.0856E-06 | -2.443 | 31.82 | ROH5;MCF2L;PRDM16;ZNF469;QSOX2;TBC1D22A;GALNT9;SLC39A3;NKX2-2;ANKS18;ENTPD2;SLC12A5;TPSG1;MTUS2;KCNAB2;BAIAP2;CBFA2T3;SEPT5;KCNQ1;H |
| hsa-miR-3180-3p | 32/1475 | 2.206E-06 | 0.000299 | 1.538E-08 | 2.0856E-06 | -2.438 | 31.75 | ROH5;MCF2L;PRDM16;ZNF469;QSOX2;TBC1D22A;GALNT9;SLC39A3;NKX2-2;ANKS18;ENTPD2;SLC12A5;TPSG1;MTUS2;KCNAB2;BAIAP2;CBFA2T3;SEPT5;KCNQ1;H |
| hsa-miR-3196 | 32/1475 | 2.206E-06 | 0.000299 | 1.538E-08 | 2.0856E-06 | -2.433 | 31.68 | ROH5;MCF2L;PRDM16;ZNF469;QSOX2;TBC1D22A;GALNT9;SLC39A3;NKX2-2;ANKS18;ENTPD2;SLC12A5;TPSG1;MTUS2;KCNAB2;BAIAP2;CBFA2T3;SEPT5;KCNQ1;H |
| hsa-miR-3186-3p | 11/270 | 3.152E-05 | 0.003562 | 5.718E-06 | 0.00043072 | -1.241 | 12.86 | KCNT1;ATCAY;C1ORF159;CHRNA4;TMEM139;PRDM16;PLD6;BAIAP2;CBFA2T3;LPP;SLC39A3 |
| hsa-miR-4707-5p | 24/1113 | 5.295E-05 | 0.004906 | 1.543E-06 | 0.00017431 | -1.926 | 18.96 | ABR;RXRA;KCNT1;JPH3;ATCAY;B3GNT6;MFNG;CDH22;PRDM16;PIEZO1;HCN2;GALNT9 |
| hsa-miR-1268 | 23/1055 | 6.513E-05 | 0.004906 | 2.207E-06 | 0.00018702 | -1.882 | 18.14 | SLC39A3 |
| hsa-miR-1268b | 23/1055 | 6.513E-05 | 0.004906 | 2.207E-06 | 0.00018702 | -1.877 | 18.09 | IAP2;CBFA2T3;LPP;ATCAY;TRIM29;KCNQ1;EFCAB2;QSOX2;DLGAP2;HCN2;GALNT9;SLC39A3 |
| hsa-miR-1250 | 15/528 | 7.953E-05 | 0.005392 | 8.034E-06 | 0.0004952 | -1.347 | 12.72 | RBFOX3;CHRNA4;MTUS2;GATA4;KCNAB2;ADARB2;CBFA2T3;SEPT5;MYLK4;C16ORF82;MPPED1;MCF2L;EFCAB2;ZNF469;HCN2 |
| hsa-miR-4462 | 19/814 | 0.0001229 | 0.006944 | 7.411E-06 | 0.0004952 | -1.614 | 14.54 | GRIA1;SLC12A5;PRKCB;CHRNA4;RASL10B;GBE1;FUT10;FLT4;CACNA1B;GATA4;ADARB2;JPH3;MPPED1;MROH5;MCF2L;TBC1D22A;PLXNA4;GALNT9;SLC39A3 |
| hsa-miR-3621 | 14/487 | 0.0001222 | 0.006944 | 1.446E-05 | 0.0007815 | -1.296 | 11.67 | TOR4A;RASL10B;ATP2B2;SEPT5;ATCAY;TRIM29;MCF2L;EFCAB2;KCNQ2;PRDM16;TPA1;TBC1D22A;PLXNA4;GALNT9 |
| hsa-miR-3676 | 16/629 | 0.0001649 | 0.007987 | 1.506E-05 | 0.0007815 | -1.450 | 12.63 | MEF2A;GRIA1;RYR2;MTUS2;GATA4;ADARB2;LPP;C1ORF159;PRDM16;GSE1;BU83;DLGAP2;GPC6;TBC1D22A;PLXNA4;GALNT9 |
| hsa-miR-191 | 15/561 | 0.000155 | 0.007987 | 1.614E-05 | 0.0007815 | -1.339 | 11.75 | TCF7L2;SLC12A5;CHRNA4;IRX4;ATP2B2;KCNAB2;CBFA2T3;LPP;SEPT5;GLIS1;AJAP1;ABR;RXRA;PRDM16;PLXNA4 |
| hsa-miR-4467 | 18/803 | 0.0003049 | 0.013781 | 2.244E-05 | 0.00101408 | -1.601 | 12.96 | MEF2A;FCN1;TOR4A;SLC12A5;IRX4;CELFA4;CHD5;MTUS2;C2CD4C;PLD6;ADARB2;GLIS1;RXRA;PIEZO1;DLGAP2;HCN2;GALNT9;NKX2-2 |
| hsa-miR-3193 | 16/695 | 0.0004984 | 0.019826 | 4.896E-05 | 0.00176735 | -1.432 | 10.89 | TOR4A;CHRNA4;CACNA1B;CELFA4;C2CD4C;NPL;ADARB2;CBFA2T3;ABR;RXRA;KCNT1;JPH3;GSE1;DLGAP2;DSCAML1;TBC1D22A |
| hsa-miR-339-3p | 13/497 | 0.0005264 | 0.019826 | 7.614E-05 | 0.00222858 | -1.226 | 9.26 | ENTPD2;RBFOX3;IRX4;CHD5;C2CD4C;BAIAP2;LPP;CACNA1G;MYLK4;RXRA;PRDM16;ZNF469;HCN2 |
| hsa-miR-574-3p | 12/431 | 0.0004986 | 0.019826 | 8.217E-05 | 0.00222858 | -1.172 | 8.91 | WWTR1;RXRA;KCNT1;RASL10B;PRDM16;MTUS2;QSOX2;PLD6;CBFA2T3;SLC39A3;ANKS18;MYLK4 |
| hsa-miR-3177-3p | 16/713 | 0.000656 | 0.023411 | 6.581E-05 | 0.00210085 | -1.459 | 10.69 | CHRNA4;CACNA1B;C2CD4C;KCNAB2;BAIAP2;GLIS1;CRHR1;CAPN14;C1ORF159;B3GNT6;PIPSKL1;MCF2L;TPAL;ALDH7A1;PLXNA4;SLC39A3 |
| hsa-miR-3178 | 17/789 | 0.0007014 | 0.023776 | 6.263E-05 | 0.00210085 | -1.561 | 11.34 | TOR4A;SLC12A5;RBFOX3;RASL10B;FLT4;BAIAP2;CBFA2T3;LPP;MR2;ABR;RXRA;MCF2L;DLGAP2;TBC1D22A;PLXNA4;HCN2;GALNT9 |
| hsa-miR-564 | 22/1188 | 0.0008977 | 0.0279 | 4.579E-05 | 0.00176735 | -1.966 | 13.79 | RA;ATCAY;C1ORF159;C1ORF159;B3GNT6;PIPSKL1;PRDM16;QSOX2;DLGAP2;PLXNA4;ANKS18 |
| hsa-miR-1306 | 11/398 | 0.0009053 | 0.0279 | 0.0001748 | 0.00408772 | -1.199 | 8.40 | ATCAY;PRKCB;CHRNA4;MCF2L;PRDM16;MTUS2;DSCAML1;LPP;PLXNA4;SLC39A3;NKX2-2 |
| hsa-miR-615-3p | 13/531 | 0.0009684 | 0.028547 | 0.0001449 | 0.00353192 | -1.273 | 8.83 | MEF2A;GRIA1;UNC5A;PRKCB;FLT4;CHD5;KCNAB2;PLD6;AJAP1;RXRA;C1ORF159;MICAL2;GSE1 |
| hsa-miR-4707-3p | 21/1145 | 0.0013297 | 0.037069 | 7.995E-05 | 0.00222858 | -1.884 | 12.48 | TOR4A;RBFOX3;CHRNA4;FLT4;IRX4;C2CD4C;KCNAB2;BAIAP2;LPP;SEPT5;C1QTNF4;ATCAY;C1ORF159;TMEM139;MCF2L;KCNQ2;PRDM16;QSOX2;TPAL;TBC1D22A;HCN2 |
| hsa-miR-4665-3p | 10/356 | 0.0013669 | 0.037069 | 0.0003065 | 0.00593688 | -1.093 | 7.21 | ABR;SLC12A5;UNC5A;FLT4;KCNQ2;PRDM16;CELFA4;GSE1;DLGAP2;AJAP1 |
| hsa-miR-1908 | 29/1849 | 0.0018737 | 0.043806 | 4.953E-05 | 0.00176735 | -3.059 | 19.21 | PRDM16;DLGAP2;TBC1D22A;PLXNA4;SLC39A3;SRRM4;ENTPD2;SLC12A5;SLC6A18;RA5L10B;ATP2B2;KCNAB2;BAIAP2;CBFA2T3;SEPT5;B3GNT6;HCN2 |
| hsa-miR-663 | 29/1849 | 0.0018737 | 0.043806 | 4.953E-05 | 0.00176735 | -3.051 | 19.16 | PRDM16;DLGAP2;TBC1D22A;PLXNA4;SLC39A3;SRRM4;ENTPD2;SLC12A5;SLC6A18;RA5L10B;ATP2B2;KCNAB2;BAIAP2;CBFA2T3;SEPT5;B3GNT6;HCN2 |
| hsa-miR-3131 | 13/569 | 0.0017953 | 0.043806 | 0.0002797 | 0.00557854 | -1.195 | 7.56 | SLC12A5;CHD5;C2CD4C;ADARB2;PRLR;MYLK4;JPH3;C1ORF159;EFCAB2;DLGAP2;PLXNA4;SLC39A3;ANKS18 |
| hsa-miR-611 | 13/569 | 0.0017953 | 0.043806 | 0.0002797 | 0.00557854 | -1.187 | 7.51 | SLC12A5;CHD5;C2CD4C;ADARB2;PRLR;MYLK4;JPH3;C1ORF159;EFCAB2;DLGAP2;PLXNA4;SLC39A3;ANKS18 |
| mmu-miR-767 | 28/1784 | 0.0022365 | 0.049 | 6.817E-05 | 0.00210085 | -2.893 | 17.65 | ZNF469;GPC6;TBC1D22A;PLXNA4;MEF2A;PRKCB;RASL10B;MTUS2;ATP2B2;KCNAB2;BAIAP2;PRLR;CAPN14;DIAPH1;TPAL;XKR6 |
| hsa-miR-4638-5p | 21/1196 | 0.0022568 | 0.049 | 0.0001459 | 0.00353192 | -1.911 | 11.65 | SRRM4;RBFOX3;CHRNA4;FUT10;FLT4;CACNA1B;LPP;MYLK4;ATCAY;ALCAM;C1ORF159;MPPED1;B3GNT6;KCNQ2;GSE1;CTSD;DLGAP2;TBC1D22A;HCN2;GALNT9;SLC39A3 |
| hsa-miR-4738-5p | 17/887 | 0.0024572 | 0.049 | 0.0002486 | 0.00526688 | -1.544 | 9.28 | TOR4A;KCNAB2;ADARB2;LPP;CRHR1;RXRA;KCNT1;ATCAY;C1ORF159;MFNG;TMEM139;KCNQ2;PRDM16;DMRTB1;DLGAP2;WDFY4;GALNT9 |
| hsa-miR-4431 | 10/383 | 0.0023355 | 0.049 | 0.0005356 | 0.00864691 | -1.040 | 6.30 | TOR4A;SRRM4;CDH22;PRDM16;GSE1;ADARB2;LPP;PLXNA4;NKX2-2;AJAP1 |
| hsa-miR-937 | 6/152 | 0.0024265 | 0.049 | 0.0009714 | 0.01197923 | -0.335 | 2.02 | SLC12A5;MCF2L;PRDM16;IRX4;C2CD4C;CBFA2T3 |

SUPPLEMENTARY DATA

Supplementary Table 7: Results from enrichment in ENCODE Histone Modification 2015 database (178 unique genes from annotation to CpG identified in Model 1)

| Term | Overlap | P-value | Adjusted P-value | Old P-value | Old Adjusted P-value | Z-score | Combined Score | Genes |
|--|---------|---------|------------------|-------------|----------------------|---------|----------------|---|
| H3K27me3_ES-Bruce4_mm9 | 48/2000 | 9.6E-11 | 3.9E-08 | 9.34E-20 | 3.81E-17 | -1.75 | 40.27 | PTPRU;RZR2;CHRNA4;ONECUT3;CACNA1B;CELF4;CHD5;GATA4;ADARB2;TRHDE;GLIS1;CRHR1;CACNA1G;C1QTNF4;MRC2;IGSF21;RXRA;JPH3;ALCAM;MROH5;CDH22;PRDM16;DMRTB1;PHACTR3;MICALL2;DLGAP2;PLXNA4;NKX2-2; ANKS1B;DIO3OS;SLC12A5;UNC5A;RBFOX3;PRKCB;RASL10B;CCDC109B;IRX4;ATP2B2;TACR1;BAIAP2;SEPT5;AJAP1;FAM101B;MPPED1;KCNQ1;KCNQ2;DSCAML1;HCN2 |
| H3K27me3_C2C12_mm9 | 34/1269 | 6.5E-09 | 1.3E-06 | 3.02E-15 | 6.16E-13 | -1.75 | 33.09 | PTPRU;RZR2;CHRNA4;ONECUT3;FLT4;CELF4;DMKN;CRHR1;CACNA1G;C1QTNF4;IGSF21;JPH3;MROH5;MCF2L;CDH22;DMRTB1;BAI1;DLGAP2;PLXNA4;GALNT9;SLC12A5;UNC5A;RBFOX3;RASL10B;ATP2B2;MIR200B;TACR1;CBFA2T3;GFRA4;PSMB9;MPPED1;KCNQ1;KCNQ2;HCN2 |
| H3K27me3_myocyte_mm9 | 28/1183 | 2E-06 | 0.00027 | 2.12E-11 | 1.44E-09 | -1.72 | 22.53 | PTPRU;RZR2;CHRNA4;ONECUT3;C2CD4C;DMKN;GLIS1;CRHR1;CACNA1G;C1QTNF4;KCNQ1;MUC2;JPH3;MROH5;MCF2L;CDH22;BAI1;DLGAP2;PLXNA4;GALNT9;NKX2-2;ENTPD2;RBFOX3;RASL10B;CBFA2T3;GFRA4;MPPED1;HCN2 |
| H3K27me3_small intestine_mm9 | 38/2000 | 5.1E-06 | 0.00052 | 2E-12 | 1.81E-10 | -1.61 | 19.67 | PTPRU;RZR2;CHRNA4;ONECUT3;CACNA1B;CHD5;GATA4;C2CD4C;NPL;TRHDE;CRHR1;C1QTNF4;IGSF21;KCNQ1;JPH3;MROH5;PRDM16;PHACTR3;BAI1;DLGAP2;PLXNA4;GALNT9;NKX2-2;ANKS1B;WWTR1;SLC12A5;UNC5A;RBFOX3;PRKCB;RASL10B; IRX4;ATP2B2;AJAP1;MPPED1;KCNQ2;ANKRD60;DSCAML1;HCN2 |
| H3K27me3_kidney epithelial cell_hg19 | 40/2217 | 9.5E-06 | 0.00077 | 2.22E-12 | 1.81E-10 | -1.65 | 19.08 | GRIA1;OCA2;RZR2;CHRNA4;CPNE4;AIRE;FLT4;CACNA1B;CHD5;GATA4;C2CD4C;NPL;ADARB2;CACNA1G;C1QTNF4;IGSF21;KCNQ1;JPH3;ATCAY;C10ORF71;MCF2L;PRDM16;PHACTR3;BAI1;PLXNA4;GALNT9;NKX2-2;ANKS1B;SRRM4;ENTPD2;SPHKAP;TPSG1; PRKCB;KCNIP4;IRX4;TACR1;GFRA4;AJAP1;C16ORF82;HAR1B |
| H3K27me3_kidney_mm9 | 32/1675 | 3E-05 | 0.00169 | 1.41E-10 | 7.19E-09 | -1.68 | 17.54 | PTPRU;RZR2;ONECUT3;SCHIP1;CACNA1B;CELF4;CHD5;GATA4;C2CD4C;TRHDE;CRHR1;CACNA1G;C1QTNF4;IGSF21;JPH3;CDH22;PRDM16;BAI1;DLGAP2;NKX2-2;DIO3OS;SRRM4;RBFOX3;PRKCB;CCDC109B;KCNIP4;MTUS2;GFRA4;KCNQ2;ANKRD60;DSCAML1;HCN2 |
| H3K27me3_bronchial epithelial cell_hg19 | 37/2082 | 3.1E-05 | 0.00169 | 2.84E-11 | 1.66E-09 | -1.61 | 16.67 | GRIA1;OCA2;RZR2;CPNE4;ONECUT3;FLT4;CACNA1B;CELF4;CHD5;C2CD4C;ADARB2;TRHDE;CACNA1G;C1QTNF4;IGSF21;JPH3;ATCAY;C10ORF71;MCF2L;CDH22;PHACTR3;GPC6;PLXNA4;ANKS1B;SRRM4;SPHKAP;PRKCB;IRX4;ATP2B2;TACR1;PRLR;GFRA4;AJAP1;MPPED1;KCNQ2;GPR123;DSCAML1 |
| H3K27me3_endothelial cell of umbilical vein_hg19 | 52/3393 | 3.3E-05 | 0.00169 | 1.98E-13 | 2.69E-11 | -1.53 | 15.76 | PTPRU;RZR2;ONECUT3;FLT4;CELF4;CHD5;DMKN;MYLK4;C1QTNF4;KCNQ1;JPH3;ATCAY;OR2AE1;MCF2L;PHACTR3;BAI1;GALNT9;ANKS1B;SRRM4;ENTPD2;UNC5A;PRKCB;AZGP1P1;TACR1;PRLR;OR1N2;GFRA4;MPPED1;KCNQ1;KCNQ2;GPR123;DSCAML1;GRIA1;OCA2;CHRNA4;AIRE;CACNA1B;C2CD4C;NPL;TRHDE;GLIS1;IGSF21;CDH22;PRDM16;GPC6;NKX2-2;RASL10B;KCNIP4;IRX4 ;ATP2B2;HCN2;HAR1B |
| H3K27me3_spleen_mm9 | 34/2000 | 0.00016 | 0.00745 | 6.82E-10 | 3.09E-08 | -1.57 | 13.66 | PTPRU;CHRNA4;FUT10;GATA4;C2CD4C;NPL;PLD6;TRHDE;CRHR1;C1QTNF4;MRC2;IGSF21;KCNQ1;ATCAY;ALCAM;MROH5;CDH22;MICALL2;GALNT9;NKX2-2;ANKS1B;WWTR1;ENTPD2;RBFOX3;RASL10B;MTUS2;ATP2B2;SEPT5;KCNQ1;KCNQ2; ANKRD60;DSCAML1;HCN2;XKR6 |
| H3K27me3_BJ_hg19 | 34/2090 | 0.00037 | 0.01527 | 2.08E-09 | 8.47E-08 | -1.53 | 12.09 | OCA2;RZR2;CPNE4;FLT4;CELF4;CHD5;C2CD4C;CACNA1G;C1QTNF4;IGSF21;ABR;JPH3;ATCAY;MCF2L;CDH22;PRDM16;RPL38;PLXNA4;GALNT9;NKX2-2;ANKS1B;ENTPD2;UNC5A;PRKCB;KCNIP4;ATP2B2;TACR1;PRLR;GFRA4;AJAP1;MPPED1;KCNQ1;KCNQ2;GPR123 |

SUPPLEMENTARY DATA

| Supplementary Table 8: Results from enrichment in GWAS Catalog 2019 database (178 unique genes from annotation to CpG identified in Model 1) | | | | | | | | |
|--|---------|----------|------------------|-------------|----------------------|---------|----------------|----------------------------|
| Term | Overlap | P-value | Adjusted P-value | Old P-value | Old Adjusted P-value | Z-score | Combined Score | Genes |
| DNA methylation (parent-of-origin) | 3-8 | 3.76E-05 | 0.017 | 4.48E-05 | 0.020 | -4.55 | 46.41 | H19;MIR675;SEPT5 |
| End-stage renal disease | 2-7 | 0.001606 | 0.207 | 0.001515 | 0.119 | -3.41 | 21.91 | MBNL1;DLGAP2 |
| Aging (time to event) | 2-8 | 0.002129 | 0.207 | 0.001886 | 0.119 | -3.42 | 21.01 | GRIA1;ATCAY |
| Disc degeneration (lumbar) | 2-9 | 0.002721 | 0.207 | 0.002295 | 0.119 | -2.81 | 16.62 | GPC6;PSMB9 |
| Asparaginase hypersensitivity in acute lymphoblastic leukemia | 2-9 | 0.002721 | 0.207 | 0.002295 | 0.119 | -2.61 | 15.40 | GRIA1;CACNA1B |
| Hair color | 4-61 | 0.002132 | 0.207 | 0.000905 | 0.119 | -1.30 | 7.99 | OCA2;PIEZO1;ZNF469;CBFA2T3 |
| Fasting blood insulin | 2-11 | 0.004109 | 0.219 | 0.003227 | 0.130 | -2.39 | 13.12 | TCF7L2;CNTN6 |
| Joint mobility (Beighton score) | 3-35 | 0.00368 | 0.219 | 0.002013 | 0.119 | -1.72 | 9.62 | PIEZO1;CRHR1;XKR6 |
| Hemoglobin A1c levels | 3-37 | 0.004312 | 0.219 | 0.002335 | 0.119 | -1.58 | 8.62 | TCF7L2;KCNQ1;PIEZO1 |
| Cutaneous squamous cell carcinoma | 2-14 | 0.006681 | 0.253 | 0.004901 | 0.132 | -2.66 | 13.33 | OCA2;LPP |
| | | | | | | | | |

SUPPLEMENTARY DATA

| Supplementary Table 9: Results from enrichment in dbGAP (accessed 08/2019) database (178 unique genes from annotation to CpG identified in Model 1) | | | | | | | | |
|---|---------|----------|------------------|-------------|----------------------|---------|----------------|---|
| Term | Overlap | P-value | Adjusted P-value | Old P-value | Old Adjusted P-value | Z-score | Combined Score | Genes |
| Body Height | 14/385 | 9.49E-06 | 0.001 | 0.0002 | 0.02702 | -2.36 | 27.29 | GRIA1;RZR2;PRKCB;CACNA1B;TRHDE;IGSF21;ATIC;BPIFA3;WDR60;CDH22;BAI1;QSOX2;ANKRD60;XKR6 |
| Body Mass Index | 13/437 | 0.000153 | 0.008 | 0.0026 | 0.11918 | -2.45 | 21.52 | RZR2;MBNL1;RBFOX3;CPNE4;FUT10;USP34;KCNIP4;ADARB2;LPP;GLIS1;KCNT1;WDFY4;ANKS1B |
| Triglycerides | 9/244 | 0.000348 | 0.013 | 0.0033 | 0.11918 | -2.29 | 18.22 | RBFOX3;CPNE4;AIRE;GBE1;KCNIP4;DLGAP2;DSCAML1;LPP;XKR6 |
| Blood Pressure | 12/454 | 0.000786 | 0.022 | 0.0095 | 0.14918 | -2.30 | 16.44 | RZR2;MPPED1;BPIFA3;CNTN6;KCNIP4;PHACTR3;MTUS2;ATP2B2;DSCAML1;GPC6;PRLR;PLXNA4 |
| Electrocardiography | 5/87 | 0.001082 | 0.024 | 0.005 | 0.13639 | -1.96 | 13.37 | RZR2;CPNE4;KCNQ1;ANKS1B;HLA-DPB2 |
| Arthritis, Rheumatoid | 4/59 | 0.001885 | 0.035 | 0.0069 | 0.14918 | -1.80 | 11.28 | PRKCB;AIRE;PHACTR3;TRHDE |
| Blood Pressure Determination | 4/64 | 0.002543 | 0.040 | 0.009 | 0.14918 | -1.88 | 11.24 | TCF7L2;KCNIP4;ADARB2;PLXNA4 |
| Echocardiography | 8/273 | 0.003158 | 0.040 | 0.0197 | 0.19685 | -1.97 | 11.36 | CNTN6;CPNE4;SCHIP1;KCNIP4;ADARB2;GPC6;PLXNA4;GLIS1 |
| Body Weight | 7/216 | 0.003272 | 0.040 | 0.0178 | 0.19616 | -1.97 | 11.25 | RBFOX3;BPIFA3;CPNE4;WDR60;FUT10;USP34;LPP |
| Elbow | 4/72 | 0.003898 | 0.043 | 0.0133 | 0.18237 | -1.68 | 9.30 | CCDC109B;PRDM16;KCNIP4;ADARB2 |
| | | | | | | | | |

SUPPLEMENTARY DATA

Supplementary Table 10: Results from enrichment in BioCarta 2016 database (178 unique genes from annotation to CpG identified in Model 1)

| Term | Overlap | P-value | Adjusted P-value | Old P-value | Old Adjusted P-value | Z-score | Combined Score | Genes |
|--|---------|---------|------------------|-------------|----------------------|---------|----------------|-----------------|
| Role of PPAR-gamma Coactivators in Obesity and Thermogenesis_Homo sapiens_h_ppargPathway | 2/9 | 0.003 | 0.158 | 0.004 | 0.256 | -1.58 | 9.33 | MED1;RXRA |
| Mechanism of Gene Regulation by Peroxisome Proliferators via PPARa_Homo sapiens_h_pparaPathway | 3/52 | 0.011 | 0.285 | 0.013 | 0.314 | -1.49 | 6.70 | MED1;RXRA;PRKCB |
| Control of Gene Expression by Vitamin D Receptor_Homo sapiens_h_vdrPathway | 2/27 | 0.024 | 0.285 | 0.030 | 0.314 | -1.39 | 5.20 | MED1;RXRA |
| VEGF, Hypoxia, and Angiogenesis_Homo sapiens_h_vegfPathway | 2/30 | 0.029 | 0.285 | 0.036 | 0.314 | -1.55 | 5.48 | PRKCB;FLT4 |
| Phospholipids as signalling intermediaries_Homo sapiens_h_edg1Pathway | 2/33 | 0.035 | 0.285 | 0.042 | 0.314 | -1.56 | 5.23 | SPHKAP;PRKCB |
| Angiotensin II mediated activation of JNK Pathway via Pyk2 dependent signaling_Homo sapiens_h_At1rPathway | 2/34 | 0.037 | 0.285 | 0.044 | 0.314 | -1.69 | 5.58 | MEF2A;PRKCB |
| Basic mechanism of action of PPARa, PPARb(d) and PPARg and effects on gene expression_Homo sapiens_h_pparPathway | 1/5 | 0.044 | 0.285 | 0.056 | 0.314 | 1.76 | -5.52 | RXRA |
| Actions of Nitric Oxide in the Heart_Homo sapiens_h_no1Pathway | 2/42 | 0.054 | 0.285 | 0.064 | 0.314 | -1.14 | 3.33 | RYR2;FLT4 |
| Signal Dependent Regulation of Myogenesis by Corepressor MITR_Homo sapiens_h_MITRPathway | 1/8 | 0.069 | 0.285 | 0.083 | 0.314 | 0.71 | -1.89 | MEF2A |
| CBL mediated ligand-induced downregulation of EGF receptors_Homo sapiens_h_cblPathway | 1/8 | 0.069 | 0.285 | 0.083 | 0.314 | 0.77 | -2.07 | PRKCB |
| | | | | | | | | |

SUPPLEMENTARY DATA

| Supplementary Table 11: Results from enrichment in WikiPathways 2019 database (178 unique genes from annotation to CpG identified in Model 1) | | | | | | | | |
|---|---------|---------|------------------|-------------|----------------------|---------|----------------|-----------------------------------|
| Term | Overlap | P-value | Adjusted P-value | Old P-value | Old Adjusted P-value | Z-score | Combined Score | Genes |
| Adipogenesis WP236 | 6/130 | 0.001 | 0.150 | 0.001 | 0.172 | -1.08 | 7.37 | MEF2A;WWTR1;MBNL1;RXRA;GATA4;PRLR |
| Genes targeted by miRNAs in adipocytes WP1992 | 2/13 | 0.006 | 0.263 | 0.008 | 0.338 | -2.98 | 15.39 | KCNQ1;HCN2 |
| Serotonin and anxiety-related events WP3944 | 2/13 | 0.006 | 0.263 | 0.008 | 0.338 | -2.91 | 15.00 | PRKCB;CRHR1 |
| Energy Metabolism WP1541 | 3/47 | 0.008 | 0.289 | 0.010 | 0.338 | -2.11 | 10.10 | MEF2A;MED1;RXRA |
| Pathways Regulating Hippo Signaling WP4540 | 4/98 | 0.011 | 0.311 | 0.013 | 0.338 | -1.22 | 5.47 | TCF7L2;PRKCB;FLT4;CDH22 |
| Sudden Infant Death Syndrome (SIDS) Susceptibility Pathways WP706 | 5/158 | 0.014 | 0.311 | 0.015 | 0.338 | -1.03 | 4.44 | RYR2;CHRNA4;KCNQ1;TACR1;NKX2-2 |
| Physiological and Pathological Hypertrophy of the Heart WP1528 | 2/25 | 0.021 | 0.404 | 0.024 | 0.468 | -2.10 | 8.14 | PRKCB;GATA4 |
| miRs in Muscle Cell Differentiation WP2012 | 2/29 | 0.027 | 0.416 | 0.031 | 0.472 | -2.48 | 8.92 | MEF2A;PRKCB |
| Non-genomic actions of 1,25 dihydroxyvitamin D3 WP4341 | 3/71 | 0.025 | 0.416 | 0.028 | 0.472 | -1.09 | 4.02 | MED1;RXRA;PRKCB |
| Cerebral Organic Acidurias, including diseases WP4519 | 1/7 | 0.061 | 0.513 | 0.069 | 0.529 | -3.50 | 9.81 | ALDH7A1 |
| | | | | | | | | |