

Additional files

Additional Table 1 Description of the genotyped polymorphisms.

| Polymorphism | Position | MAF | Alleles (M/m) | HWE P | Call rate (%) |
|---------------------|-----------------|------------|----------------------|--------------|----------------------|
| rs9939609 | Intron 1 | 0.444 | T/A | 0.682 | 99.4 |
| rs9928094 | Intron 1 | 0.462 | A/G | 0.789 | 99.8 |
| rs9930333 | Intron 1 | 0.463 | T/G | 0.687 | 99.4 |
| rs9935401 | Intron 1 | 0.439 | G/A | 1 | 99.3 |
| rs10852521 | Intron 1 | 0.446 | C/T | 0.606 | 99.8 |
| rs1861868 | Intron 1 | 0.466 | A/G | 0.797 | 99.8 |
| rs7205986 | Intron 1 | 0.084 | A/G | 0.246 | 99.8 |
| rs11643744 | Intron 1 | 0.267 | A/G | 0.256 | 99.8 |
| rs7203521 | Intron 1 | 0.383 | A/G | 1 | 100 |
| rs12445162 | Intron 1 | 0.075 | G/A | 0.681 | 100 |
| rs11075986 | Intron 1 | 0.084 | C/G | 0.201 | 100 |
| rs17818902 | Intron 3 | 0.245 | T/G | 0.855 | 99.8 |
| rs3826169 | Intron 3 | 0.228 | T/C | 1 | 100 |
| rs8061518 | Intron 3 | 0.309 | A/G | 0.889 | 97.9 |
| rs7190053 | Intron 3 | 0.183 | C/T | 1 | 99.1 |
| rs2111114 | Intron 4 | 0.212 | A/G | 0.843 | 99.8 |
| rs8044353 | Intron 4 | 0.053 | G/A | 1 | 99.8 |
| rs10521303 | Intron 5 | 0.439 | C/A | 0.069 | 99.1 |
| rs1558756 | Intron 6 | 0.444 | C/T | 0.119 | 100 |
| rs16952623 | Intron 6 | 0.132 | T/C | 0.388 | 99.4 |
| rs7194336 | Intron 7 | 0.441 | G/T | 0.199 | 100 |

| | | | | | |
|------------|----------|-------|-----|--------------|------|
| rs9929152 | Intron 7 | 0.277 | A/G | 0.040 | 99.6 |
| rs16952649 | Intron 7 | 0.122 | C/T | 0.008 | 100 |
| rs10852525 | Intron 7 | 0.106 | G/A | 0.159 | 99.1 |
| rs8053966 | Intron 7 | 0.184 | T/C | 0.007 | 99.6 |
| rs2111113 | Intron 7 | 0.074 | G/C | 0.330 | 99.6 |
| rs16952624 | Exon 7 | 0.001 | C/T | 1 | 99.6 |
| rs1136002 | Intron 8 | 0.317 | T/C | 0.462 | 100 |
| rs2075202 | Intron 8 | 0.025 | T/G | 1 | 100 |
| rs2540781 | Intron 8 | 0.128 | C/A | 0.345 | 100 |
| rs12932373 | Intron 8 | 0.138 | C/T | 0.589 | 99.4 |
| rs1008400 | Intron 8 | 0.459 | C/T | 0.153 | 99.6 |
| rs1125337 | Intron 8 | 0.450 | A/T | 0.027 | 99.8 |
| rs17833492 | Intron 8 | 0.313 | C/A | 0.466 | 99.6 |
| rs7203181 | Intron 8 | 0.339 | C/A | 0.780 | 100 |
| rs7194907 | Intron 8 | 0.479 | T/C | 0.091 | 98.9 |
| rs2689248 | Intron 8 | 0.480 | C/A | 0.368 | 99.4 |
| rs13334214 | Intron 8 | 0.201 | C/T | 0.228 | 98.3 |
| rs1558687 | Intron 8 | 0.242 | C/T | 0.864 | 99.4 |
| rs4784351 | Intron 8 | 0.270 | A/G | 0.742 | 97.6 |
| rs12596638 | Intron 8 | 0.136 | G/A | 1 | 99.8 |
| rs8056299 | Intron 8 | 0.429 | A/G | 0.696 | 99.8 |
| rs7194243 | Intron 8 | 0.219 | C/T | 0.337 | 100 |
| rs6499656 | Intron 8 | 0.130 | G/C | 0.356 | 100 |
| rs7199716 | Intron 8 | 0.394 | C/T | 0.893 | 99.8 |
| rs8054364 | Intron 8 | 0.146 | C/G | 0.040 | 99.8 |

| | | | | | |
|------------|----------|-------|-----|-------|------|
| rs8049235 | Intron 8 | 0.393 | G/A | 0.285 | 99.6 |
| rs17225435 | Intron 8 | 0.115 | A/G | 1 | 100 |
| rs7200579 | Intron 8 | 0.060 | C/G | 0.536 | 100 |
| rs697771 | Intron 8 | 0.427 | C/T | 0.362 | 100 |
| rs7191513 | Intron 8 | 0.424 | G/A | 0.241 | 100 |
| rs8049933 | Intron 8 | 0.108 | C/T | 0.284 | 99.3 |

HWE P: Hardy-Weinberg equilibrium (p for the control group); FAM: minor allele frequency.

Additional Table 2 Association of rs9928094 with biomarkers of inflammation and cardiovascular disease risk in obese children.

| | AA | AG | GG | β (95%CI) | P |
|--|--------------|--------------|--------------|----------------------|--------------|
| n | 155 | 263 | 115 | | |
| Anthropometric factors | | | | | |
| Height (m) | 1.38 ± 0.01 | 1.39 ± 0.01 | 1.39 ± 0.01 | -0.003 (-0.01, 0.00) | 0.167 |
| Weight (kg) | 42.2 ± 1.4 | 45.9 ± 1.1 | 48.8 ± 2.0 | 3.14 (1.37, 4.91) | 0.001 |
| BMI (kg/m ²) | 21.55 ± 0.49 | 23.02 ± 0.36 | 24.12 ± 0.61 | 1.26 (0.53, 1.98) | 0.001 |
| BMI z-score | 1.43 ± 0.17 | 1.88 ± 0.13 | 2.25 ± 0.20 | 0.44 (0.18, 0.69) | 0.001 |
| Waist circumference (cm) | 70.43 ± 1.40 | 73.16 ± 1.01 | 76.45 ± 1.61 | 2.77 (0.03, 5.51) | 0.048 |
| Clinical and metabolic biomarkers | | | | | |
| Systolic BP (mm Hg) | 104 ± 1 | 105 ± 1 | 105 ± 1 | 0.90 (-0.91, 2.72) | 0.330 |
| Diastolic BP (mm Hg) | 63 ± 1 | 65 ± 1 | 65 ± 1 | 0.81 (-0.67, 2.28) | 0.285 |
| Glucose (mg/dl) | 85 ± 1 | 84 ± 0 | 84 ± 1 | -0.37 (-1.22, 0.49) | 0.399 |
| Insulin (mU/l) | 8.81 ± 0.66 | 8.86 ± 0.43 | 9.35 ± 0.65 | 0.02 (-0.02, 0.05) | 0.349 |
| HOMA-IR | 1.88 ± 0.15 | 1.87 ± 0.10 | 1.98 ± 0.14 | 0.01 (-0.02, 0.05) | 0.443 |
| Triacylglycerol (mg/dl) | 68 ± 3 | 63 ± 2 | 68 ± 3 | -0.64 (-0.45, 3.27) | 0.748 |

| | | | | | |
|---|---------------|---------------|---------------|----------------------|--------------|
| Apo-AI (mg/dl) | 143 ± 2 | 140 ± 2 | 136 ± 3 | -3.41 (-6.72, -0.09) | 0.044 |
| Cholesterol (mg/dl) | 169 ± 3 | 166 ± 2 | 168 ± 2 | -0.001 (-0.01, 0.01) | 0.811 |
| HDL-c (mg/dl) | 58 ± 1 | 57 ± 1 | 55 ± 1 | -0.84 (-2.65, 0.96) | 0.360 |
| Adiponectin (mg/l) | 25.94 ± 0.99 | 25.15 ± 0.73 | 24.22 ± 1.06 | -0.50 (-1.93, 0.93) | 0.496 |
| Resistin (µg/l) | 10.28 ± 0.46 | 10.91 ± 0.35 | 11.15 ± 0.51 | 0.50 (-0.19, 1.19) | 0.158 |
| Leptin (µg/l) | 13.69 ± 1.29 | 14.50 ± 0.84 | 15.55 ± 1.38 | 1.00 (-0.78, 2.78) | 0.273 |
| Inflammatory biomarkers | | | | | |
| C-reactive protein (mg/l) | 2.02 ± 0.30 | 2.31 ± 0.24 | 2.59 ± 0.48 | 0.31 (-0.19, 0.81) | 0.228 |
| IL-6 (ng/l) | 5.73 ± 0.93 | 5.66 ± 0.62 | 6.67 ± 1.21 | 0.53 (-0.83, 1.89) | 0.448 |
| IL-8 (ng/l) | 1.87 ± 0.16 | 1.90 ± 0.14 | 1.95 ± 0.22 | 0.06 (-0.21, 0.34) | 0.645 |
| TNFα (ng/l) | 3.58 ± 0.17 | 3.69 ± 0.13 | 3.30 ± 0.18 | -0.07 (-0.32, 0.18) | 0.568 |
| Cardiovascular disease risk biomarkers | | | | | |
| MMP-9 (µg/l) | 82.36 ± 4.57 | 80.95 ± 3.67 | 94.46 ± 5.61 | 0.04 (0.003, 0.07) | 0.034 |
| MPO (µg/l) | 17.77 ± 2.16 | 17.67 ± 1.61 | 17.63 ± 1.75 | 0.14 (-2.91, 3.19) | 0.929 |
| sE-Selectin (µg/l) | 26.06 ± 1.17 | 28.43 ± 1.06 | 26.38 ± 1.33 | 0.61 (-1.27, 2.50) | 0.525 |
| sICAM-1 (mg/l) | 0.162 ± 0.005 | 0.168 ± 0.005 | 0.160 ± 0.005 | 0.004 (-0.01, 0.01) | 0.993 |

| | | | | | |
|---------------------|--------------|--------------|--------------|--------------------|-------|
| Active PAI-1 (μg/l) | 8.39 ± 0.60 | 8.64 ± 0.52 | 10.04 ± 0.88 | 0.80 (-0.22, 1.82) | 0.127 |
| Total PAI-1 (μg/l) | 23.04 ± 1.34 | 22.22 ± 0.94 | 26.72 ± 1.90 | 0.02 (-0.01, 0.06) | 0.223 |

CI: Confidence interval; BMI: body mass index; BP: blood pressure; HOMA-IR: homeostasis model assessment for insulin resistance; HDL-c: high-density lipoprotein cholesterol; IL: interleukin; TNF-α: tumor necrosis factor alpha; MMP-9: metalloproteinase-9; MPO: myeloperoxidase; sICAM-1: soluble intracellular adhesion molecule-1, PAI-1: plasminogen activator inhibitor. β Coefficients represent the change in absolute traits values of each additional risk allele. General linear models were used to examine associations, P adjusted by age and sex.

Additional Table 3 Association of rs9930333 with biomarkers of inflammation and cardiovascular disease risk in obese children.

| | TT | TG | GG | β (95% CI) | P |
|--|--------------|--------------|--------------|----------------------|--------------|
| n | 154 | 262 | 115 | | |
| Anthropometric factors | | | | | |
| Height (m) | 1.38 ± 0.01 | 1.39 ± 0.01 | 1.39 ± 0.01 | -0.003 (-0.01, 0.00) | 0.192 |
| Weight (kg) | 42.4 ± 1.4 | 46.0 ± 1.1 | 48.5 ± 2.0 | 3.07 (1.31, 4.84) | 0.001 |
| BMI (kg/m ²) | 21.60 ± 0.49 | 23.09 ± 0.36 | 24.03 ± 0.61 | 1.23 (0.52, 1.96) | 0.001 |
| BMI z-score | 1.44 ± 0.17 | 1.90 ± 0.13 | 2.23 ± 0.20 | 0.43 (0.18, 0.69) | 0.001 |
| Waist circumference (cm) | 70.54 ± 1.41 | 73.31 ± 1.02 | 76.30 ± 1.61 | 2.73 (-0.003, 5.47) | 0.051 |
| Clinical and metabolic biomarkers | | | | | |
| Systolic BP (mm Hg) | 104 ± 1 | 105 ± 1 | 105 ± 1 | 0.88 (-0.93, 2.69) | 0.339 |
| Diastolic BP (mm Hg) | 63 ± 1 | 65 ± 1 | 65 ± 1 | 0.83 (-0.65, 2.30) | 0.272 |
| Glucose (mg/dl) | 85 ± 1 | 84 ± 0 | 84 ± 1 | -0.37 (-1.22, 0.49) | 0.402 |
| Insulin (mU/l) | 8.83 ± 0.67 | 8.89 ± 0.43 | 9.31 ± 0.65 | 0.02 (-0.02, 0.05) | 0.400 |
| HOMA-IR | 1.89 ± 0.15 | 1.88 ± 0.10 | 1.97 ± 0.14 | 0.01 (-0.02, 0.05) | 0.499 |
| Triacylglycerols (mg/dl) | 68 ± 3 | 63 ± 2 | 68 ± 3 | -0.42 (-4.32, 3.48) | 0.034 |

| | | | | | |
|---|---------------|---------------|---------------|----------------------|--------------|
| Apo-AI (mg/dl) | 143 ± 2 | 141 ± 2 | 135 ± 3 | -3.59 (-6.90, -0.29) | 0.033 |
| Cholesterol (mg/dl) | 169 ± 3 | 167 ± 2 | 168 ± 2 | -0.002 (-0.01, 0.01) | 0.728 |
| HDL-c (mg/dl) | 57 ± 1 | 57 ± 1 | 55 ± 1 | -0.97 (-2.77, 0.83) | 0.293 |
| Adiponectin (mg/l) | 25.77 ± 0.98 | 25.37 ± 0.75 | 23.78 ± 1.01 | -0.73 (-2.16, 0.70) | 0.319 |
| Resistin (µg/l) | 10.28 ± 0.46 | 10.95 ± 0.35 | 11.10 ± 0.51 | 0.45 (-0.23, 1.14) | 0.198 |
| Leptin (µg/l) | 13.76 ± 1.29 | 14.56 ± 0.84 | 15.56 ± 1.38 | 0.98 (-0.80, 2.75) | 0.283 |
| Inflammatory biomarkers | | | | | |
| C-reactive protein (mg/l) | 2.03 ± 0.30 | 2.31 ± 0.24 | 2.60 ± 0.48 | 0.32 (-0.18, 0.82) | 0.371 |
| IL-6 (ng/l) | 5.74 ± 0.93 | 5.67 ± 0.62 | 6.68 ± 1.21 | 0.51 (-0.84, 1.87) | 0.461 |
| IL-8 (ng/l) | 1.86 ± 0.16 | 1.90 ± 0.15 | 1.95 ± 0.22 | 0.06 (-0.21, 0.33) | 0.656 |
| TNFα (ng/l) | 3.57 ± 0.17 | 3.70 ± 0.13 | 3.32 ± 0.18 | -0.08 (-0.32, 0.17) | 0.552 |
| Cardiovascular disease risk biomarkers | | | | | |
| MMP-9 (µg/l) | 81.61 ± 4.54 | 80.95 ± 3.71 | 96.34 ± 5.60 | 0.04 (0.006, 0.07) | 0.021 |
| MPO (µg/l) | 17.65 ± 2.17 | 17.78 ± 1.62 | 17.57 ± 1.75 | 0.07 (-2.97, 3.12) | 0.962 |
| sE-Selectin (µg/l) | 26.06 ± 1.17 | 28.43 ± 1.06 | 26.38 ± 1.33 | 0.56 (-1.33, 2.44) | 0.562 |
| sICAM-1 (mg/l) | 0.162 ± 0.005 | 0.169 ± 0.005 | 0.159 ± 0.005 | -0.000 (-0.01, 0.01) | 0.953 |

| | | | | | |
|---------------------|--------------|--------------|--------------|--------------------|-------|
| Active PAI-1 (µg/l) | 8.40 ± 0.60 | 8.69 ± 0.52 | 10.06 ± 0.88 | 0.79 (-0.23, 1.81) | 0.129 |
| Total PAI-1 (µg/l) | 23.02 ± 1.35 | 22.13 ± 0.94 | 27.04 ± 1.90 | 0.03 (-0.01, 0.06) | 0.187 |

CI: Confidence interval; BMI: body mass index; BP: blood pressure; HOMA-IR: homeostasis model assessment for insulin resistance; HDL-c: high-density lipoprotein cholesterol; IL: interleukin; TNF- α : tumor necrosis factor alpha; MMP-9: metalloproteinase-9; MPO: myeloperoxidase; sICAM-1: soluble intracellular adhesion molecule-1, PAI-1: plasminogen activator inhibitor. β Coefficients represent the change in absolute traits values of each additional risk allele. General linear models were used to examine associations, P adjusted by age and sex.

Additional Table 4 Association of rs9935401 with biomarkers of inflammation and cardiovascular disease risk in obese children.

| | GG | AG | AA | β (95%CI) | P |
|--|--------------|--------------|--------------|-----------------------|------------------|
| n | 167 | 261 | 102 | | |
| Anthropometric factors | | | | | |
| Height (m) | 1.38 ± 0.01 | 1.39 ± 0.01 | 1.40 ± 0.02 | -0.003 (-0.01, 0.001) | 0.115 |
| Weight (kg) | 42.1 ± 1.4 | 45.9 ± 1.1 | 50.1 ± 2.2 | 3.44 (1.66, 5.22) | <0.001 |
| BMI (kg/m ²) | 21.56 ± 0.47 | 23.08 ± 0.37 | 24.51 ± 0.64 | 1.37 (0.65, 2.10) | <0.001 |
| BMI z-score | 1.42 ± 0.16 | 1.93 ± 0.13 | 2.32 ± 0.22 | 0.47 (0.22, 0.73) | <0.001 |
| Waist circumference (cm) | 70.29 ± 1.35 | 73.35 ± 1.01 | 77.46 ± 1.71 | 3.37 (0.62, 6.14) | 0.017 |
| Clinical and metabolic biomarkers | | | | | |
| Systolic BP (mm Hg) | 104 ± 1 | 105 ± 1 | 107 ± 1 | 1.19 (-0.65, 3.02) | 0.205 |
| Diastolic BP (mm Hg) | 63 ± 1 | 65 ± 1 | 66 ± 1 | 1.39 (-0.10, 2.88) | 0.068 |
| Glucose (mg/dl) | 85 ± 1 | 84 ± 0 | 84 ± 1 | -0.42 (-1.28, 0.44) | 0.342 |
| Insulin (mU/l) | 8.78 ± 0.62 | 8.78 ± 0.44 | 9.78 ± 0.71 | 0.02 (-0.01, 0.05) | 0.209 |
| HOMA-IR | 66 ± 3 | 64 ± 2 | 68 ± 4 | 0.34 (-3.61, 4.29) | 0.866 |
| Triacylglycerols (mg/dl) | 1.88 ± 0.14 | 1.86 ± 0.10 | 2.07 ± 0.16 | 0.02 (-0.02, 0.05) | 0.286 |

| | | | | | |
|---|---------------|---------------|---------------|----------------------|-------|
| Apo-AI (mg/dl) | 143 ± 2 | 141 ± 2 | 135 ± | -3.04 (-6.38, 0.31) | 0.076 |
| Cholesterol (mg/dl) | 168 ± 2 | 166 ± 2 | 170 ± 2 | 0.003 (-0.01, 0.01) | 0.584 |
| HDL-c (mg/dl) | 58 ± 1 | 57 ± 1 | 55 ± 1 | -1.04 (-2.86, 0.79) | 0.266 |
| Adiponectin (mg/l) | 26.09 ± 0.95 | 24.80 ± 0.73 | 24.16 ± 1.10 | -0.67 (-2.11, 0.78) | 0.366 |
| Resistin (µg/l) | 10.30 ± 0.43 | 10.85 ± 0.36 | 11.43 ± 0.55 | 0.59 (-0.10, 1.29) | 0.096 |
| Leptin (µg/l) | 13.29 ± 1.20 | 14.68 ± 0.84 | 16.29 ± 1.52 | 1.51 (-0.28, 3.30) | 0.100 |
| Inflammatory biomarkers | | | | | |
| C-reactive protein (mg/l) | 2.05 ± 0.29 | 2.29 ± 0.23 | 2.77 ± 0.53 | 0.38 (-0.12, 1.89) | 0.140 |
| IL-6 (ng/l) | 5.56 ± 0.87 | 6.00 ± 0.70 | 6.19 ± 1.05 | 0.44 (-0.93, 1.82) | 0.528 |
| IL-8 (ng/l) | 1.86 ± 0.15 | 1.96 ± 0.15 | 1.83 ± 0.21 | 0.03 (-0.24, 0.31) | 0.819 |
| TNFα (ng/l) | 3.52 ± 0.16 | 3.69 ± 0.13 | 3.33 ± 0.20 | -0.01 (-0.26, 0.24) | 0.962 |
| Cardiovascular disease risk biomarkers | | | | | |
| MMP-9 (µg/l) | 82.52 ± 4.42 | 82.49 ± 3.73 | 92.82 ± 6.05 | 0.03 (-0.004, 0.07) | 0.080 |
| MPO (µg/l) | 17.65 ± 2.02 | 17.92 ± 1.63 | 17.65 ± 1.88 | 0.27 (-2.81, 3.35) | 0.865 |
| sE-Selectin (µg/l) | 26.18 ± 1.12 | 28.25 ± 1.07 | 26.64 ± 1.40 | 0.74 (-1.16, 2.64) | 0.448 |
| sICAM-1 (mg/l) | 0.162 ± 0.005 | 0.169 ± 0.005 | 0.158 ± 0.006 | 0.0003 (-0.01, 0.01) | 0.951 |

| | | | | | |
|---------------------|--------------|--------------|--------------|--------------------|-------|
| Active PAI-1 (µg/l) | 8.41 ± 0.58 | 8.62 ± 0.52 | 10.35 ± 0.97 | 0.90 (-0.13, 1.93) | 0.089 |
| Total PAI-1 (µg/l) | 23.26 ± 1.31 | 22.18 ± 0.92 | 26.84 ± 2.09 | 0.02 (-0.02, 0.06) | 0.313 |

CI: Confidence interval; BMI: body mass index; BP: blood pressure; HOMA-IR: homeostasis model assessment for insulin resistance; HDL-c: high-density lipoprotein cholesterol; IL: interleukin; TNF- α : tumor necrosis factor alpha; MMP-9: metalloproteinase-9; MPO: myeloperoxidase; sICAM-1: soluble intracellular adhesion molecule-1, PAI-1: plasminogen activator inhibitor. β Coefficients represent the change in absolute traits values of each additional risk allele. General linear models were used to examine associations, P adjusted by age and sex.