

**Supplementary Table 3.** Median with IQR values of sCD40 and sCD40L in plasma (A.U., SUMMIT cohort) when comparing between two groups (Mann-Whitney U test)

|                              | CD40 (n=1,438)                         |                      | CD40L (n=1,438)                       |                     |
|------------------------------|--|----------------------|---------------------------------------|---------------------|
|                              | Content (A.U.)                         | <i>P</i> , adjusted* | Content (A.U.)                        | <i>P</i> , adjusted |
| Female sex (vs. male sex)    | 9.16 (8.71–9.90) vs. 9.14 (8.70–9.89)  | 0.94                 | 6.37 (5.04–8.06) vs. 6.99 (5.62–8.24) | 0.0016              |
| Diabetes type 2 (yes vs. no) | 9.18 (8.74–9.91) vs. 9.08 (8.61–9.84)  | 0.016                | 6.94 (5.48–8.27) vs. 6.52 (5.17–8.02) | 0.020               |
| Stroke (yes vs. no)          | 9.75 (9.08–10.37) vs. 9.12 (8.70–9.80) | 0.000030             | 7.88 (5.62–8.65) vs. 6.75 (5.37–8.16) | 0.020               |
| Hypertension (yes vs. no)    | 9.24 (8.73–9.95) vs. 8.98 (8.37–9.76)  | 0.00021              | 6.86 (5.39–8.21) vs. 6.63 (5.33–8.18) | 0.51                |
| AMI (yes vs. no)             | 9.27 (8.70–9.98) vs. 9.10 (8.70–9.82)  | 0.016                | 7.13 (5.47–8.24) vs. 6.71 (5.34–8.19) | 0.28                |

IQR, interquartile range; sCD40, soluble CD40; sCD40L, soluble CD40 ligand; A.U., arbitrary unit; SUMMIT, The SURrogate markers for Micro- and Macro-vascular hard endpoints for Innovative diabetes Tools; AMI, acute myocardial infarction.

\**P*-value adjusted for multiple comparisons (according to the Holm-Šidák test).

**Supplementary Table 4.** Spearman correlations between the circulating levels of sCD40 and sCD40L (A.U., SUMMIT cohort) and change in IMT in the CCA and the carotid bulb region assessed by vascular ultrasound (at the 3-year follow-up point compared to study inclusion)

| IMT change (mm)  | CD40                                       |          |                      | CD40L                                      |                        |                        |
|------------------|--|----------|----------------------|--|------------------------|------------------------|
|                  | (n=1,438 at inclusion; n=798 at follow-up) |          |                      | (n=1,438 at inclusion; n=798 at follow-up) |                        |                        |
|                  | <i>r</i>                                   | <i>P</i> | <i>P</i> , adjusted* | <i>r</i>                                   | <i>P</i>               | <i>P</i> , adjusted    |
| <b>Right</b>     |  |          |                      |  |                        |                        |
| Mean IMT in CCA  | -0.046                                     | 0.18     | 0.63                 | 0.014                                      | 0.69                   | 0.90                   |
| Max IMT in CCA   | 0.015                                      | 0.67     | 0.82                 | 0.090                                      | 0.009                  | 0.027                  |
| Mean IMT in bulb | 0.027                                      | 0.44     | 0.82                 | 0.149                                      | 0.000022               | 0.00011                |
| Max IMT in bulb  | 0.075                                      | 0.034    | 0.21                 | 0.243                                      | 3.65×10 <sup>-12</sup> | 2.55×10 <sup>-11</sup> |
| <b>Left</b>      |  |          |                      |  |                        |                        |
| Mean IMT in CCA  | -0.039                                     | 0.25     | 0.68                 | 0.014                                      | 0.69                   | 0.90                   |
| Max IMT in CCA   | 0.051                                      | 0.14     | 0.59                 | 0.115                                      | 0.001                  | 0.0040                 |
| Mean IMT in bulb | 0.025                                      | 0.47     | 0.82                 | 0.233                                      | 1.66×10 <sup>-10</sup> | 9.96×10 <sup>-10</sup> |
| Max IMT in bulb  | 0.092                                      | 0.009    | 0.070                | 0.261                                      | 7.86×10 <sup>-14</sup> | 6.29×10 <sup>-13</sup> |

sCD40, soluble CD40; sCD40L, soluble CD40 ligand; A.U., arbitrary unit; SUMMIT, The SURrogate markers for Micro- and Macro-vascular hard endpoints for Innovative diabetes Tools; IMT, intima-media thickness; CCA, common carotid artery.

\**P*-value adjusted for multiple comparisons.