

SUPPLEMENTARY DATA

Supplementary Table 1. Clinical characteristics of 224 participants with diabetes, according to retinopathy status

| Characteristics | All participants N=224 | Retinopathy | | P-value* |
|-------------------------------------|---------------------------|------------------|--------------------|----------|
| | | Absent N = 80 | Present N = 144 | |
| (Person Data) | | | | |
| Gender, % male | 59.4 | 55.8 | 62.0 | 0.35 |
| Current cigarette smoker, % yes | 7.1 | 58.1 | 50.4 | 0.11 |
| Use of Insulin, % | 65.6 | 57.9 | 71.3 | 0.037 |
| Use of oral hypoglycemic agents, % | 54.9 | 53.7 | 55.8 | 0.75 |
| Use of lipid medication, % | 54.5 | 46.3 | 58.1 | 0.080 |
| Albuminuria, % | | | | |
| Micro | 2.7 | 2.2 | 3.1 | 0.42 |
| Macro | 12.5 | 5.9 | 17.1 | <0.001 |
| Diabetes type, % type 1 | 38.0 | 38.9 | 37.2 | 0.79 |
| Diabetic retinopathy severity, % | | | | |
| No DR | 35.7 | | | |
| Mild NPDR | 14.3 | - | - | - |
| Moderate NPDR | 24.5 | - | - | - |
| Severe NPDR - PDR | 25.5 | - | - | - |
| Mean (SD)/ Median (IQR) | | | | |
| †Age, years | 59 (15) | 58 (18) | 60 (14) | 0.43 |
| BMI, kg/m ² | 30.4 (6.3) | 29.5 (5.6) | 31.1 (6.6) | 0.070 |
| Systolic BP, mmHg | 128.0 (14.5) | 125.2 (13.1) | 130.1 (15.1) | 0.011 |
| †Duration of diabetes, years | 15 (12) | 10 (11) | 18 (13) | <0.001 |
| HbA _{1c} , % | 7.9 (1.4) | 7.6 (1.7) | 8.0 (1.2) | 0.022 |
| Total Cholesterol, mmol/L | 4.6 (1.1) | 4.7 (1.1) | 4.5 (1.1) | 0.47 |
| HDL-C, mmol/L | 1.4 (0.4) | 1.5 (0.4) | 1.3 (0.5) | 0.056 |
| Non HDL-C, mmol/L | 3.2 (1.0) | 3.3 (1.0) | 3.2 (1.1) | 0.85 |
| LDL-C, mmol/L | 2.5 (0.8) | 2.5 (0.7) | 2.5 (0.9) | 0.99 |
| †Triglyceride, mmol/L | 1.3 (1.0) | 1.3 (1.0) | 1.3 (1.1) | 0.50 |
| Apolipoprotein AI, g/L | 1.4 (0.3) | 1.5 (0.2) | 1.4 (0.3) | 0.001 |
| Apolipoprotein B, g/L | 0.8 (0.3) | 0.8 (0.2) | 0.9 (0.3) | 0.015 |
| Apolipoprotein B/AI | 0.6 (0.2) | 0.6 (0.2) | 0.7 (0.2) | <0.001 |
| †Glomerular Filtration Rate, mL/min | 77 (61 – 91) | 76 (61 – 91) | 77 (61 – 90) | 0.090 |

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| | | | | |
|--|-------------------|--------------------|--------------------|----------|
| †Albumin-creatinine ratio, mg/mmol | 1.06 (0.57 – 2.8) | 1.00 (0.51 – 2.30) | 1.10 (0.60 – 3.73) | 0.16 |
| ‡Skin iontophoresis, % | | | | |
| Sodium nitroprusside (SNP) | 5.0 (2.3) | 5.6 (2.6) | 4.6 (2.1) | 0.005 |
| Acetylcholine (ACh) | 6.4 (3.2) | 7.2 (3.1) | 5.9 (3.1) | 0.007 |
| | | | | |
| | All eyes | | Retinopathy | P-value* |
| | | Absent | Present | |
| (Eye-specific data) | N= 420 | N = 186 | N = 234 | |
| ‡Diabetic retinopathy severity, % | | | | |
| No DR | 41.6 | | | |
| Mild NPDR | 13.4 | - | - | - |
| Moderate NPDR | 22.3 | - | - | - |
| Severe NPDR - PDR | 22.7 | - | - | - |
| Without laser | 16.0 | - | - | - |
| With laser | 6.7 | - | - | - |
| Grid/ Focal laser | 2.6 | - | - | - |
| Pan Retinal Photocoagulation | 4.1 | - | - | - |
| ‡Flicker-light induced vasodilatation, % | | | | |
| Arteriolar dilatation | 1.4 (2.1) | 1.8 (2.1) | 1.2 (1.9) | 0.006 |
| Venular dilatation | 2.8 (2.1) | 3.2 (1.8) | 2.6 (2.5) | 0.003 |
| ‡Retinal vascular tortuosity index ($\times 10^5$) | | | | |
| Arteriolar | 34.4 (18.6) | 33.6 (16.8) | 37.2 (21.9) | 0.043 |
| Venular | 39.7 (18.8) | 39.1 (19.2) | 40.2 (18.5) | 0.57 |
| ‡Artery to Vein ratio (AVR) | 0.67 (0.07) | 0.69 (0.06) | 0.67 (0.07) | 0.13 |

*P-values were obtained using chi-square test (categorical), t-test (continuous and normally distributed) or Mann-Whitney U test (continuous and skewed), comparing diabetes participants with and without retinopathy. †Results are shown as median (first – third quartile). ‡Right and left side data were used. Analysis using cluster t-test, with person as clustering factor to account within-person correlation between right and left side.

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Supplementary Table 2. Associations between serum traditional lipids and skin iontophoresis and flicker-light induced vasodilatation

| Serum Traditional Lipids | Skin iontophoresis (%)* | | | | Flicker-light induced vasodilatation (%)* | | | |
|-----------------------------|--------------------------|---------|--------------------------|---------|---|---------|--------------------------|---------|
| | Sodium nitroprussid | | Acetylcholine | | Arteriolar vasodilatation | | Venular vasodilatation | |
| | Mean change ^a | P value | Mean change ^a | P value | Mean change ^a | P value | Mean change ^a | P value |
| (per SD increase) | (95% CI) | | (95% CI) | | (95% CI) | | (95% CI) | |
| Cholesterol, (1.1 mmol/L) | -0.14 (-0.66, 0.37) | 0.58 | 0.51 (-0.04, 1.06) | 0.07 | 0.07 (-0.14, 0.28) | 0.52 | 0.04 (-0.18, -0.25) | 0.73 |
| HDL, (0.4 mmol/L) | 0.68 (0.00, 1.35) | 0.049 | 1.27 (0.57, 1.98) | <0.001 | 0.30 (-0.05, 0.66) | 0.069 | 0.00 (-0.28, 0.28) | 0.99 |
| Non HDL, (0.2 mmol/L) | -0.29 (-0.95, 0.37) | 0.40 | -0.34 (-1.06, 0.38) | 0.36 | -0.16 (-0.42, 0.09) | 0.20 | 0.08 (-0.19, 0.36) | 0.55 |
| LDL, (0.3 mmol/L) | 0.04 (-0.63, 0.74) | 0.92 | -0.21 (-0.97, 0.56) | 0.60 | -0.07 (-0.33, 0.19) | 0.59 | 0.07 (-0.22, 0.36) | 0.64 |
| Triglycerides, (1.0 mmol/L) | -0.52 (-1.08, 0.04) | 0.069 | -0.31 (-0.92, 0.29) | 0.31 | 0.02 (-0.21, 0.26) | 0.86 | -0.08 (-0.31, 0.16) | 0.54 |

* Percentage changes relative to baseline measures

^a Data are age and gender adjusted mean change

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Supplementary Table 3. Associations between serum traditional lipids and retinal vascular tortuosity index

| Serum Traditional Lipids | Retinal Vascular Tortuosity index ($\times 10^5$) | | | |
|---------------------------------|---|---------------------|--------------------------------------|---------|
| | Arteriolar | | Venuilar | |
| | Mean change ^a (per SD increase) | P value (95% CI) | Mean change ^a (95% CI) | P value |
| Total-Cholesterol, (1.1 mmol/L) | -0.39 (-2.57, 1.80) | 0.73 | -1.51 (-3.11, 0.09) | 0.07 |
| HDL-C, (0.4 mmol/L) | 0.03 (-3.14, 3.21) | 0.98 | 1.58 (-0.37, 3.53) | 0.11 |
| Non HDL, (0.2 mmol/L) | -0.81 (-3.72, 2.11) | 0.59 | -2.33 (-5.11, 0.54) | 0.10 |
| LDL-C, (0.3 mmol/L) | -0.84 (-3.82, 2.15) | 0.58 | -2.79 (-5.62, 0.97) | 0.10 |
| Triglycerides, (1.0 mmol/L) | -1.48 (-3.78, 0.81) | 0.21 | 0.31 (-1.37, 1.99) | 0.72 |

* P-value for trend

^a Data are age and gender adjusted mean change