

## Supplementary Data

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# Metabolic Syndrome, Diabetes, Poor Cognition, and Dementia in the Caerphilly Prospective Study

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Table 1  
Regression coefficients \* (and 95% CI) for metabolic syndrome (NCEP ATP3 definition) † not diabetes, diabetes ‡ and cognitive outcomes at phase 5

| Exposure                            | Model | Clinical outcomes: odds ratio<br>(95% confidence interval) |                   | Neuropsychological tests: beta coefficients<br>(95% confidence interval) |                     |
|-------------------------------------|-------|--|-------------------|--|---------------------|
|                                     |       | CIND   | Dementia          | AH4  | CAMCOG              |
| <b>Phase 1 (n = 995)</b>            |       |  |                   |  |                     |
| Metabolic syndrome† (not diabetes‡) | 1     | 0.94 (0.53, 1.67)  | 0.62 (0.26, 1.53) | -0.78 (-2.86, 1.30)  | 0.73 (-1.74, 3.21)  |
|                                     | 2     | 0.87 (0.43, 1.77)  | 0.74 (0.29, 1.89) | 0.02 (-1.60, 1.64)   | 0.32 (-1.75, 2.39)  |
| Diabetes‡ (±metabolic syndrome †)   | 1     | 5.14 (2.21, 11.95)   | 1.19 (0.24, 5.75) | -5.61 (-10.08, -1.14)  | -2.69 (-7.82, 2.44) |
|                                     | 2     | 4.04 (1.40, 11.63)   | 0.59 (0.07, 5.22) | -4.26 (-7.83, -0.69)   | 0.04 (-4.49, 4.57)  |
| <b>Phase 2 (n = 1131)</b>           |       |  |                   |  |                     |
| Metabolic syndrome† (not diabetes‡) | 1     | 1.18 (0.81, 1.73)  | 1.59 (0.97, 2.60) | -1.98 (-3.36, -0.59)   | -0.39 (-2.04, 1.27) |
|                                     | 2     | 1.05 (0.69, 1.61)  | 1.54 (0.89, 2.66) | -0.75 (-1.80, 0.31)  | 0.05 (-1.41, 1.51)  |
| Diabetes‡ (±metabolic syndrome†)    | 1     | 2.18 (1.17, 4.06)  | 1.10 (0.40, 3.05) | -5.85 (-8.75, -2.95)   | -1.77 (-5.11, 1.58) |
|                                     | 2     | 1.63 (0.80, 3.35)  | 0.74 (0.23, 2.39) | -2.76 (-5.00, -0.51)   | -0.05 (-3.06, 2.95) |
| <b>Phase 3 (n = 1109)</b>           |       |  |                   |  |                     |
| Metabolic syndrome† (not diabetes‡) | 1     | 0.97 (0.67, 1.41)  | 1.24 (0.76, 2.02) | -0.26 (-1.59, 1.07)  | -0.59 (-2.14, 0.96) |
|                                     | 2     | 0.86 (0.57, 1.30)  | 1.15 (0.67, 1.99) | 0.73 (-0.29, 1.74)   | 0.34 (-1.05, 1.74)  |
| Diabetes‡ (±metabolic syndrome†)    | 1     | 1.19 (0.65, 2.19)  | 1.03 (0.44, 2.39) | -3.86 (-6.29, -1.43)   | -2.51 (-5.26, 0.24) |
|                                     | 2     | 0.84 (0.41, 1.69)  | 0.86 (0.34, 2.20) | -2.02 (-3.85, -0.19)   | -0.62 (-3.08, 1.85) |

\*Model 1 adjusted for age at exposure used continuously. Model 2 additionally adjusted for confounders (social class, smoking, alcohol, C3 NART). Max n given. In both models reference group is normal (neither metabolic syndrome nor diabetes); †according to [2]; ‡According to self report or [24].

Table 2  
Regression coefficients (and 95% CI) for individual components of metabolic syndrome and cognitive outcomes at phase 5 in men without diabetes\*

| Exposure                             | Clinical outcomes: odds ratio<br>(95% confidence interval) |                    | Neuropsychological tests: beta coefficients<br>(95% confidence interval) |                      |
|--------------------------------------|--|--------------------|--|----------------------|
|                                      | CIND   | Dementia           | AH4  | CAMCOG               |
| <b>Phase 1 (n = 779)†</b>            |  |                    |  |                      |
| Systolic blood pressure (10 mmHg)‡   | 0.82 (0.67, 1.00)  | 1.06 (0.85, 1.31)  | -0.42 (-0.86, 0.02)  | -0.20 (-0.76, 0.36)  |
| Diastolic blood pressure (10 mmHg)‡  | 1.17 (0.86, 1.58)  | 1.03 (0.73, 1.45)  | -0.29 (-0.95, 0.38)  | -0.01 (-0.87, 0.85)  |
| Fasting plasma glucose (mmol/L)      | 1.99 (0.23, 16.92)   | 0.68 (0.05, 9.06)  | -0.95 (-5.97, 4.07)  | -0.06 (-6.63, 6.51)  |
| Fasting triglycerides (mmol/L)       | 1.18 (0.74, 1.87)  | 0.75 (0.42, 1.33)  | 0.29 (-0.80, 1.37)   | -0.52 (-1.95, 0.90)  |
| Fasting HDL cholesterol (mmol/L)     | 0.55 (0.27, 1.12)  | 0.79 (0.33, 1.88)  | 1.48 (-0.08, 3.04)   | 0.95 (-1.1, 3.01)    |
| Body Mass Index (kg/m <sup>2</sup> ) | 0.97 (0.9, 1.05)   | 0.96 (0.86, 1.07)  | 0.02 (-0.17, 0.21)   | 0.03 (-0.21, 0.28)   |
| <b>Phase 2 (n = 965)†</b>            |  |                    |  |                      |
| Systolic blood pressure (10 mmHg)‡   | 1.02 (0.89, 1.16)  | 1.09 (0.92, 1.29)  | -0.18 (-0.5, 0.15)   | -0.53 (-0.96, -0.10) |
| Diastolic blood pressure (10 mmHg)‡  | 1.12 (0.87, 1.43)  | 0.97 (0.70, 1.34)  | -0.17 (-0.77, 0.44)  | 0.44 (-0.37, 1.24)   |
| Fasting plasma glucose (mmol/L)      | 1.60 (0.18, 14.21)   | 1.23 (0.08, 19.46) | -0.82 (-6.23, 4.59)  | 3.17 (-3.94, 10.28)  |
| Fasting triglycerides (mmol/L)       | 0.88 (0.56, 1.39)  | 1.08 (0.60, 1.94)  | -0.01 (-1.13, 1.11)  | 0.60 (-0.87, 2.08)   |
| Fasting HDL cholesterol (mmol/L)     | 0.37 (0.15, 0.97)  | 0.68 (0.21, 2.25)  | 0.65 (-1.61, 2.92)   | 0.61 (-2.42, 3.64)   |
| Body Mass Index (kg/m <sup>2</sup> ) | 0.97 (0.91, 1.04)  | 0.96 (0.87, 1.05)  | 0.03 (-0.14, 0.20)   | 0.00 (-0.23, 0.22)   |
| <b>Phase 3 (n = 924)†</b>            |  |                    |  |                      |
| Systolic blood pressure (10 mmHg)‡   | 1.05 (0.92, 1.19)  | 1.20 (1.01, 1.42)  | -0.40 (-0.72, -0.08)   | -0.55 (-0.98, -0.13) |
| Diastolic blood pressure (10 mmHg)‡  | 0.92 (0.72, 1.16)  | 0.86 (0.63, 1.19)  | 0.36 (-0.22, 0.94)   | 0.74 (-0.04, 1.53)   |
| Fasting plasma glucose (mmol/L)      | 4.98 (0.58, 42.84)   | 2.39 (0.15, 39.07) | -1.98 (-7.21, 3.24)  | -0.98 (-7.98, 6.02)  |
| Fasting triglycerides (mmol/L)       | 1.22 (0.81, 1.83)  | 0.98 (0.56, 1.71)  | 0.11 (-0.92, 1.15)   | -0.02 (-1.40, 1.36)  |
| Body Mass Index (kg/m <sup>2</sup> ) | 0.98 (0.91, 1.05)  | 0.95 (0.87, 1.05)  | 0.01 (-0.15, 0.18)   | 0.14 (-0.08, 0.37)   |
| <b>Average values (n = 1031)</b>     |  |                    |  |                      |
| Systolic blood pressure (10 mmHg)‡   | 1.00 (0.85, 1.17)  | 1.20 (0.97, 1.48)  | -0.37 (-0.77, 0.02)  | -0.88 (-1.42, -0.34) |
| Diastolic blood pressure (10 mmHg)‡  | 1.05 (0.77, 1.42)  | 0.88 (0.60, 1.30)  | -0.18 (-0.91, 0.54)  | 0.82 (-0.19, 1.82)   |
| Fasting plasma glucose (mmol/L)      | 0.94 (0.75, 1.18)  | 0.84 (0.61, 1.15)  | -0.28 (-0.88, 0.32)  | 0.00 (-0.79, 0.79)   |
| Fasting triglycerides (mmol/L)       | 1.08 (0.89, 1.31)  | 1.00 (0.74, 1.36)  | 0.00 (-0.5, 0.51)  | 0.05 (-0.65, 0.74)   |
| Fasting HDL cholesterol (mmol/L)     | 0.47 (0.18, 1.24)  | 0.90 (0.26, 3.13)  | -0.16 (-2.46, 2.14)  | -0.14 (-3.35, 3.06)  |
| Body Mass Index (kg/m <sup>2</sup> ) | 0.98 (0.92, 1.04)  | 0.98 (0.89, 1.07)  | 0.01 (-0.15, 0.18)   | 0.06 (-0.17, 0.28)   |

\*components according to WHO definition [3]. Adjusted for age at exposure used continuously, all other components and confounders (social class, smoking, alcohol, C3 NART). Max n given. Men with diabetes self report or [24] excluded; †max n, size for each regression varies; ‡for blood pressure the coefficient is for a 10 unit increase.