Supplementary Material

Supplementary Table 1. Associations of creatinine-based estimated glomerular filtration rate with impaired fasting glucose and impaired glucose tolerance (dichotomous outcomes) among 4136 participants without diabetes in the Cardiovascular Health Study

| | | Adjusted difference (95% confidence interval) | | |
|------------------------------|----------------|---|------------------|--|
| Estimated GFR | Unadjusted | Model 1 | Model 2 | |
| (mL/min/1.73m ²) | Prevalence (%) | | | |
| Impaired Fasting Glue | cose (mg/dL) | | | |
| ≥90 | 44.3 | 1 (reference) | 1 (reference) | |
| 75-89 | 44.1 | 0.89 (0.73,1.08) | 0.90 (0.73,1.11) | |
| 60-74 | 49.2 | 1.02 (0.83,1.26) | 0.98 (0.79,1.22) | |
| 45-59 | 51.6 | 1.13 (0.90,1.42) | 1.02 (0.80,1.29) | |
| < 45 | 50.2 | 1.02 (0.75,1.40) | 0.81 (0.58,1.14) | |
| Continuous** | | 1.05 (1.02,1.09) | 1.01 (0.97,1.05) | |
| | | P=0.02 | P=0.51 | |
| Impaired Glucose Tol | | | | |
| ≥90 | 41.5 | 1 (reference) | 1 (reference) | |
| 75-89 | 42.5 | 0.96 (0.78,1.17) | 0.95 (0.77,1.17) | |
| 60-75 | 41.7 | 0.88 (0.71,1.09) | 0.83 (0.66,1.03) | |
| 45-59 | 42.9 | 0.88 (0.71,1.12) | 0.77 (0.61,0.98) | |
| < 45 | 46.8 | 0.88 (0.64,1.21) | 0.69 (0.49,0.97) | |
| Continuous** | | 0.98(0.42,1.02) | 0.94 (0.90,0.98) | |
| | | P=0.33 | P=0.006 | |

Included are 607 participants with estimated GFR \geq 90 mL/min/1.73m², 1379 participants with estimated GFR 75-89 mL/min/1.73m², 1109 participants with estimated GFR 60-74 mL/min/1.73m², 774 participants with estimated GFR 45-59 mL/min/1.73m², and 267 participants with estimated GFR < 45 mL/min/1.73m².

IFG: impaired fasting glucose (fasting glucose 100-125mg/dL)

IGT: impaired glucose tolerance (2-hour glucose 149-199mg/dL)

Model 1 is adjusted for age, sex, race, site, and education

Model 2 is additionally adjusted for leisure time physical activity, smoking, alcohol, waist circumference, dietary score, medications and prevalent cardiovascular disease.

**Continuous models evaluate eGFR models per 10 mL/min/1.73m² lower eGFR

| | | Adjusted difference* (95% confidence interval) | | |
|---|------------------------|---|-------------------|--|
| Estimated GFR (mL/min/1.73m ²) | Unadjusted Mean, SD | Model 1 | Model 2 | |
| Fasting insulin (% | 50 | | | |
| difference) | | | | |
| ≥ 90 | 12.0(6.0) | 1(reference) | 1(reference) | |
| 75-89 | 13.1(7.1) | 8.4(4.5,12.4) | 4.7(1.3,8.2) | |
| 60-74 | 14.6(7.6) | 22.5(18.0,27.2) | 10.9(7.0,14.9) | |
| 45-59 | 16.1(8.0) | 37.6(30.9,44.7) | 20.2(14.8,25.9) | |
| < 45 | 16.7(8.7) | 45.3(34.4,57.0) | 23.7(15.0,33.0) | |
| Continuous** | 10.7(8.7) | 6.0(5.3,6.8) | 3.5(2.8,4.2) | |
| Continuous | | P<0.001 | P<0.001 | |
| | | P<0.001 | P<0.001 | |
| ISI (% difference) | 1.0/2.5 | 10(6) | 1.0/ 6 | |
| ≥ 90 | 4.0(2.5) | 1.0(reference) | 1.0(reference) | |
| 75-89 | 3.8(2.3) | -4.7(-9.6,0.4) | -0.5(-5.2,-0.4) | |
| 60-74 | 3.4(2.1) | -16.7(-21.0,-12.0) | -5.3(-10.1,-0.3) | |
| 45-59 | 3.1(2.0) | -25.0(-30.2,-19.4) | -10.2(-16.1,-4.0) | |
| < 45 | 2.9(1.8) | -30.6(-37.4,-23.0) | -13.2(-21.6,-4.0) | |
| Continuous** | | -6.2(-7.4,-5.1) | -2.5(-3.6,-1.4) | |
| | | P<0.001 | P<0.001 | |
| Beta-cell function index | | | | |
| (pmol/L) | | | | |
| \geq 90 | 1264(508) | 0(reference) | 0(reference) | |
| 75-89 | 1377(572) | 130.4(82.5,178.2) | 99.9(53.7,146.0) | |
| 60-74 | 1435(636) | 217.4(164.7,270.1) | 131.4(78.6,184.1 | |
| 45-59 | 1511(665) | 332.8(260.7,404.8) | 214.5(143.7,285.4 | |
| < 45 | 1509(761) | 368.1(242.1,494.1) | 225.2(102.3,348.) | |
| Continuous** | | 43.9(33.6,54.2) | 38.5(27.3,49.7) | |
| | | P<0.001 | P<0.001 | |
| | | | 1 (01001 | |
| 2-hour glucose (mg/dl) | | | | |
| \geq 90 | 137.8(44) | 0(reference) | 0(reference) | |
| 75-89 | 133.1(40) | -5.3(-9.0,-1.2) | -6.8(-10.4,-3.1) | |
| 60-74 | 138.6(42) | -0.7(-4.6,3.2) | -5.4(-9.3,-1.4) | |
| 45-59 | 142.0(43) | 0.7(-4.2,5.7) | -6.2(-11.2,-1.1) | |
| < 45 | 153.7(46) | 9.6(1.9,17.3) | 0.8(-7.2,8.8) | |
| Continuous** | | 0.9(0.16,1.8) | -0.4(-1.3,0.4) | |

Supplementary Table 2. Associations of cystatin C-based estimated glomerular filtration rate with fasting insulin, ISI, beta cell function, and 2-hour glucose among 3638 participants without prevalent diabetes in the Cardiovascular Health Study.

Included are 897 participants with estimated GFR \geq 90 mL/min/1.73m², 1120 participants with estimated GFR 75-89 mL/min/1.73m², 1010 participants with estimated GFR 60-74 mL/min/1.73m², 448 participants with estimated GFR 45-59 mL/min/1.73m², and 163 participants with estimated GFR < 45 mL/min/1.73m².

Model 1 is adjusted for age, sex, race, site, and education

Model 2 is additionally adjusted for leisure time physical activity, smoking, alcohol, waist circumference, dietary score, medications and prevalent cardiovascular disease.

* Differences expressed in mg/dL for fasting and 2-hour glucose concentration (absolute difference), pmol/L for beta cell function (absolute difference) or as percent difference for fasting insulin concentration and ISI (relative difference).

**Continuous models evaluate eGFR models per 10 mL/min/1.73m² lower eGFR; p-values are generated using this continuous model.

Supplementary Table 3. Associations of cystatin C-based estimated glomerular filtration rate with incident diabetes among 4170 participants in the Cardiovascular Health Study.

| Estimated GFR (mL/min/1.73m ²) | Incident DM Number of events | Incidence rate | Model 1 | Model 2 |
|---|------------------------------------|-------------------|-----------------|-----------------|
| ≥ 90 | 117 | 8.8 | 1(reference) | 1(reference) |
| 75-89 | 118 | 7.6 | 0.86(0.66,1.10) | 0.75(0.58,0.97) |
| 60-74 | 110 | 9.1 | 1.06(0.82,1.37) | 0.77(0.58,1.00) |
| 45-59 | 42 | 9.7 | 1.21(0.84,1.73) | 0.80(0.54,1.17) |
| < 45 | 14 | 11.5 | 1.41(0.80,2.48) | 0.84(0.45,1.55) |
| Continuous** | | | 1.04(0.98,1.11) | 0.96(0.90,1.02) |
| | | | P=0.16 | P=0.15 |

Included are 1056 participants with estimated GFR \geq 90 mL/min/1.73m², 1295 participants with estimated GFR 75-89 mL/min/1.73m², 1146 participants with estimated GFR 60-74 mL/min/1.73m², 492participants with estimated GFR 45-59 mL/min/1.73m², and 181 participants with estimated GFR < 45 mL/min/1.73m².

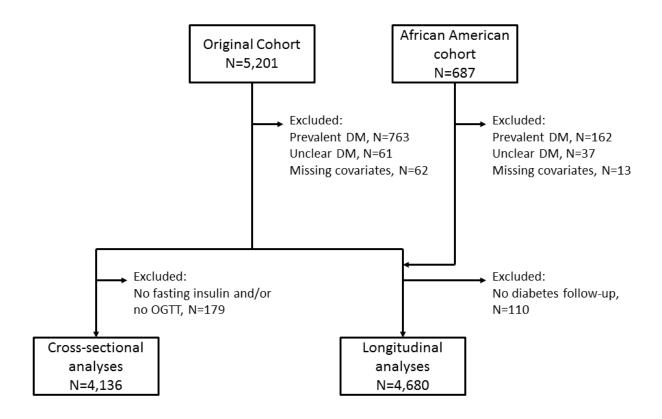
Model 1 is adjusted for age, sex, race, site, and education

Model 2 is additionally adjusted for leisure time physical activity, smoking, alcohol, waist circumference, dietary score, medications and prevalent cardiovascular disease.

**Continuous models evaluate eGFR models per 10 mL/min/1.73m² lower eGFR; p-values are generated using this continuous model.

Supplementary Figures

Supplementary Figure 1. Cardiovascular Health Study participants included in this study.



Supplementary Figure 2. Correlation of beta-cell function index (pmol/L) with insulin sensitivity index among 4136 participants in the Cardiovascular Health Study.

