| Table 1: Diabetes Complications Phenotype Definitions and Descriptions | | | | | |
|--|-----------------------|---|---------------------------|------------------------|--|
| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | |
| Type II Diabetes (T2D) | | Have you ever been diagnosed with diabetes? | Anti-diabetes medications | No T2D on all criteria | |
| ICD/OPCS-4 Codes: ICD-9 codes: 250.XX or ICD-10 codes: (E11.0–11.9) | | | | | |
| Notes: OGTT: Oral glucose tolerance test, FPG: fasting plasma glucose, HbA1C: Hemoglobin A1C | | | | | |

Self-report

Medical records

Control group

Normal AER/ACR

| | creatinine ratio (ACR) 2.5-25 mg/mmol (men); 3.5-35 mg/mmol (women) | | | | | |
|--|---|--------------|-----------------|----------------|--|--|
| ICD/OPCS-4 Codes: NA | | | | | | |
| Notes: AER: urine albumin excretion rat | Notes: AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable | | | | | |
| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | | |
| Macroalbuminuria (macro) | AER >= 200ug/min or 300mg/ 24h or ACR > 25/35 mg/mmol (men/women) | | | Normal AER/ACR | | |
| ICD/OPCS-4 Codes: NA | | | | | | |
| Notes: AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable | | | | | | |
| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | | |
| Albuminuria (ALB) | Microalbuminuria or Macroalbuminuria | | | Normal AER/ACR | | |
| ICD/OPCS-4 Codes: NA | • | - | • | • | | |
| Notes: | | | | | | |
| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | | |
| End-stage renal disease (ESRD vs. cont | trol) eGFR < 15 mL/min/1.73 m2 or dialysis or transplant | | | Normal AER/ACR | | |
| ICD/OPCS-4 Codes: ICD-10 codes: (E85.3, N16.5, N18.0, N18.5, Q60.1, T82.4, T86.1, Y60.2, Y60.2, Y61.2, Y62.2, Y84.1, Z49.0, Z49.1, Z49.2, Z94.0, Z99.2); OPCS-4 coded procedures (L74.1, L74.2, L74.3, L74.4, L74.5, L74.6, L74.8, L74.9, M01.2, | | | | | | |
| M01.3. M01.4. M01.5. M01.8. M01.9. | 01.3, M01.4, M01.5, M01.8, M01.9, M02.3, M08.4, M17.2, M17.4, M17.8, M17.9, X40.1, X40.2, X40.3, X40.4, X40.5, X40.6, X40.7, X40.8, X40.9, X41.1, X41.2, X41.8, X41.9, X42.1, X42.8, X42.9, X43.1) LIKBB participants without evidence of a CKD | | | | | |

M01.3, M01.4, M01.5, M01.8, M01.9, M02.3, M08.4, M17.2, M17.4, M17.4, M17.9, M04.0, 1, X40.2, X40.3, X40.4, X40.5, X40.6, X40.7, X40.8, X40.9, X41.1, X41.2, X41.8, X42.9, X42.1, X42.8, X42.9, X43.1) UKBB participants without evidence of a Ck stage indicator are excluded (i.e. those with acute kidney injury are excluded) refer to: https://biobank.ndph.ox.ac.uk/showcase/showcase/docs/alg_outcome_esrd.pdf

Notes: ESRD: End-stage renal disease, eGFR: estimated Glomerular filtration rate (using creatinine & Epi-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self report | Medical records | Control group |
|---------------------------|-----------------------|-------------|-----------------|----------------|
| Diabetic Nephropathy (DN) | Macro or ESRD | | | Normal AER/ACR |
| ICD/OPCS-4 Codes: NA | • | • | • | • |

Notes: eGFR: estimated Glomerular filtration rate (calculated using creatinine & EPI-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

Albumin excretion rate (AER) 20-200 ug/min or 30-300mg/24h OR Albumin-

Lab/Clinical Criteria

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|-------------------------------|--|-------------|-----------------|----------------|
| Diabetic Kidney Disease (DKD) | Microalbuminuria, Macroalbuminuria or ESRD | | | Normal AER/ACR |
| ICD/OPCS-4 Codes: NA | | | | |

Notes: eGFR: estimated Glomerular filtration rate (calculated using creatinine & EPI-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|------------------------------|--------------------------|-------------|-----------------|-----------------------|
| Chronic Kidney Disease (CKD) | eGFR < 60 mL/min/1.73 m2 | | | eGFR≥60mL/min/1.73 m2 |
| ICD/OPCS-4 Codes: NA | | | | |

Notes: eGFR: estimated Glomerular filtration rate (calculated using creatinine & EPI-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|--------|--------------------------------------|-------------|-----------------|--|
| CKD-DN | Macroalbuminuria or ESRD and eGFR<45 | | | eGFR ≥60mL/min/1.73 m2 and Normal AER/ACR |
| | • | • | = | • |

ICD/OPCS-4 Codes: NA

Microalbuminuria (micro)

Notes: eGFR: estimated Glomerular filtration rate (calculated using creatinine & EPI-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|----------------------|--|-------------|-----------------|---|
| CKD-DKD | Microalbuminuria, Macroalbuminuria or ESRD and eGFR<45 | | | eGFR≥60mL/min/1.73 m2 and Normal AER/ACR |
| ICD/OPCS-4 Codes: NA | | | | |

Votes: eGFR: estimated Glomerular filtration rate (calculated using creatinine & EPI-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|----------------|--|-------------|-----------------|------------------|
| ESRD vs. Macro | eGFR < 15 mL/min/1.73 m2 or dialysis or transplant | | | Macroalbuminuria |

ICD/OPCS-4 Codes: ICD-10 codes: (E85.3, N16.5, N18.0, N18.5, Q60.1, T82.4, T86.1, Y60.2, Y60.2, Y60.2, Y61.2, Y62.2, Y84.1, Z49.0, Z49.1, Z49.0, Z49.1, Z49.2, Z94.0, Z99.2); OPCS-4 coded procedures (L74.1, L74.2, L74.3, L74.4, L74.5, L74.6, L74.8, L74.9, M01.2, M01.3, M01.4, M01.5, M01.8, M01.9, M01.9, M02.3, M08.4, M17.2, M17.4, M17.8, M17.9, X40.1, X40.2, X40.3, X40.4, X40.5, X40.5, X40.6, X40.7, X40.8, X40.9, X41.1, X41.2, X41.8, X41.9, X42.1, X42.8, X42.9, X43.1) UKBB participants without evidence of a CKD stage indicator are excluded (i.e. those with acute kidney injury are excluded) refer to: https://biobank.ndph.ox.ac.uk/showcase/showcase/docs/alg_outcome_esrd.pdf

Notes: ESRD: End-stage renal disease, eGFR: estimated Glomerular filtration rate (using creatinine & Epi-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|------------------|--|-------------|-----------------|---------------|
| ESRD vs. No ESRD | eGFR < 15 mL/min/1.73 m2 or dialysis or transplant | | | Non-ESRD |

ICD/OPCS-4 Codes: ICD-10 codes: (E85.3, N16.5, N18.0, N18.5, Q60.1, T82.4, T86.1, Y60.2, Y60.2, Y61.2, Y62.2, Y84.1, Z49.0, Z49.1, Z49.2, Z94.0, Z99.2); OPCS-4 coded procedures (L74.1, L74.2, L74.3, L74.4, L74.5, L74.6, L74.8, L74.9, M01.2, M01.3, M01.4, M01.5, M01.8, M01.9, M01.9, M02.3, M08.4, M17.2, M17.4, M17.8, M17.9, X40.1, X40.2, X40.3, X40.4, X40.5, X40.6, X40.7, X40.8, X40.9, X41.1, X41.2, X41.8, X41.9, X42.1, X42.8, X42.9, X43.1) UKBB participants without evidence of a CKD stage indicator are excluded (i.e. those with acute kidney injury are excluded) refer to: https://biobank.ndph.ox.ac.uk/showcase/showcase/docs/alg_outcome_esrd.pdf

Notes: ESRD: End-stage renal disease, eGFR: estimated Glomerular filtration rate (using creatinine & Epi-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|-------------------------|---|-------------|-----------------|------------------------|
| CKD-Extreme vs eGFR>=60 | eGFR < 15 mL/min/1.73 m2 or dialysis or transplant | | | eGFR≥60mL/min/1.73 m2 |
| (ESRD vs. CKD-Control) | eGFR < 15 mL/min/ 1.73 m2 or dialysis or transplant | | | eGFR 260mL/min/1./3 m2 |

ICD/OPCS-4 Codes: ICD-10 codes: (E85.3, N16.5, N18.0, N18.5, Q60.1, T82.4, T86.1, Y60.2, Y60.2, Y61.2, Y62.2, Y84.1, Z49.0, Z49.1, Z49.2, Z94.0, Z99.2); OPCS-4 coded procedures (L74.1, L74.2, L74.3, L74.4, L74.5, L74.6, L74.8, L74.9, M01.2, M01.3, M01.4, M01.5, M01.8, M01.9, M01.9, M02.3, M08.4, M17.2, M17.4, M17.8, M17.9, X40.1, X40.2, X40.3, X40.4, X40.5, X40.5, X40.6, X40.7, X40.8, X40.9, X41.1, X41.2, X41.8, X41.9, X42.1, X42.8, X42.9, X43.1) UKBB participants without evidence of a CKC stage indicator are excluded (i.e. those with acute kidney injury are excluded) refer to: https://biobank.ndph.ox.ac.uk/showcase/showcase/docs/alg_outcome_esrd.pdf

Notes: ESRD: End-stage renal disease, eGFR: estimated Glomerular filtration rate (using creatinine & Epi-CKD formula), AER: urine albumin excretion rate, ACR: urine albumin-creatinine ratio, NA: Not applicable

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | |
|--|--|-------------------------------------|-----------------------|------------------------------|--|
| | Fundus exam: Early Treatment in Diabetic Retinopathy Study (ETDRS) 1. Absent/ present | Hx of retinopathy Ex. Have you been | | No Hx of retinopathy | |
| Diabetic Retinopathy (DRET) | | told that you have diabetic eye | ICD code: Retinopathy | Absent retinopathy on fundus | |
| | | disease? | | exam | |
| ICD/OPCS-4 Codes: ICD-9 codes (362.0); ICD-10 code: (H36.0, E10.3, E11.3, E12.3, E13.3, E14.3) | | | | | |
| Notes: ETDRS Early Treatment in Diabetic Retinopathy Study | | | | | |

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | |
|--|--|-------------|---|-------------------------|--|
| Non-alcoholic fatty liver disease (NAFLD) | Abdominal CT-scan (Liver/Spleen attenuation ratio < 1) | | ICD code: Non-alcoholic fatty liver disease | No evidence of NAFLD on | |
| | | | | imaging | |
| | | | | Normal AST/ALT | |
| | | | | No NAFLD diagnosis | |
| ICD/OPCS-4 Codes: ICD-9 codes (571.5, 571.8, 571.9); ICD_10 codes (K75.8, K76.0, K74.0, K74.6, K76.9) | | | | | |
| Notes: AST: Aspartate transaminase, ALT: alanine aminotransferase, significant alcohol consumption is defined as >21 standard drinks per week in men and >14 standard drinks per week in women over a 2-year period preceding baseline liver | | | | | |

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | |
|---|--|----------------|-------------------------------------|------------------------|--|
| Peripheral artery disease (PAD) | Ankle brachial index (ABI) values < 0.9 or Intermittent claudication | History of PAD | ICD code: Peripheral artery disease | ABI betwen 1.0 and 1.4 | |
| ICD/OPCS-4 Codes: ICD-9 codes: (440.0, 440.2, 443.8, 443.9); ICD-10 codes (I70.0, 170.2, 170.8, 170.9, 173.8, 173.9); OPCS-4 coded procedures (X09.3, X09.4, X09.5, 121.6, L51.3, L51.6, L51.8, L52.1, L52.2, L54.1, L54.4, L54.8, L59.1, L59.2, L59.3, | | | | | |
| Notes: ABI: Ankle brachial index | | | | | |

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group | |
|--|---|--|------------------------------------|----------------------------|--|
| Coronary Heart Disease (CHD) | Myocardial infarction (MI) or revascularization procedure | History MI or revascularization or CHD | ICD9/ICD10 MI or revascularization | No hx CHD for all criteria | |
| ICD/OPCS-4 Codes: ICD-9 codes: (410-412.9, 414) or OPCS-4 coded procedure: (K40-K49-K50, and K75); ICD-10 codes: (121-125.9) | | | | | |
| Notes: revasc: revascularization, MI: Myocardial infarction, CHD: coronary heart disease | | | | | |

| Trait | Lab/Clinical Criteria | Self-report Self-report | Medical records | Control group |
|---|--------------------------------|-------------------------|-------------------|-------------------------------|
| STROKE | Any Stroke (not including TIA) | History of stroke | ICD9/ICD10 stroke | No hx stroke for all criteria |
| ICD/OPCS-4 Codes: ICD-9 codes: (430, 431, 434, 436); ICD-10 codes: (160-64) | | | | |
| Notes: TIA: Transient ischemic attack | | | | |

| Trait | Lab/Clinical Criteria | Self-report | Medical records | Control group |
|--|--|----------------|--------------------------------------|----------------------------|
| Cardiovascular Disease (CVD) | CHD or STROKE or AFIB or Heart Failure | History of CVD | ICD9/ICD10 CVD (Angina not included) | No hx CVD for all criteria |
| ICD/OPCS-4 Codes: Heart Failure ICD-9 codes: (428.0, 428.9); ICD-10 codes: (ICD-10 codes: ICD-10 codes: (ICD-10 codes: ICD-10 codes: IC | | | | |
| (160-64) or ICD-9 codes: (430, 431, 434, 436) | | | | |
| Notes: CHD: coronary heart disease, AFIB: atrial fibrillation, CVD: Cardiovascular Disease | | | | |