Supplemental Appendices

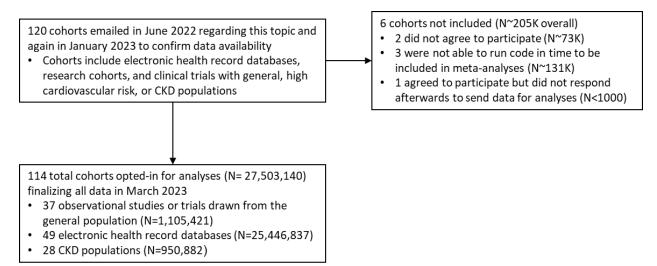
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Appendix 1. Data analysis overview and analytic notes for some individual cohorts

1.1 Overview:

In order to determine the cohorts eligible for this paper, in June 2022 all cohorts in CKD-PC were contacted regarding their interest in this topic and in January 2023 cohorts were contacted to confirm availability of data.



As previously described,¹ the collaborating cohorts were asked to compile a dataset with approximately 30 variables (main exposure [serum creatinine to estimate GFR, albuminuria, age, sex, race/ethnicity, history of cardiovascular diseases, smoking, diabetes, diabetes medications systolic blood pressure, antihypertensive medications, total cholesterol, HDL cholesterol, use of statins, BMI], outcome [all-cause mortality, cardiovascular mortality, myocardial infarction, stroke, heart failure, peripheral artery disease, atrial fibrillation, kidney failure with replacement therapy, acute kidney injury, hospitalizations]).

To be consistent across cohorts, the CKD-PC Data Coordinating Center sent definitions for those medical history variables to participating cohorts (outlined below). We instructed studies not to impute any variables and let us know of any differences in definitions.

Medical history variable	Definition
History of cardiovascular disease	Prior diagnosis of CVD based on myocardial infarction, bypass grafting,
(CVD)	percutaneous coronary intervention, heart failure, or stroke. Identification
	at any time during study period (from baseline to follow-up).
History of coronary heart disease	Prior diagnosis of CHD based on myocardial infarction, bypass grafting,
(CHD)	percutaneous coronary intervention. Identification at any time during study
	period (from baseline to follow-up).
History of heart failure (HF)	Prior diagnosis of heart failure. Identification at any time during study
Thistory of heart failure (Th')	period (from baseline to follow-up).
History of stroke	Prior diagnosis of stroke. Identification at any time during study period
	(from baseline to follow-up).
History of atrial fibrillation (Afib)	Prior diagnosis of atrial fibrillation. Identification at any time during study
Thistory of atrial hormation (And)	period (from baseline to follow-up).
History of peripheral artery disease	Prior diagnosis of peripheral artery disease. Identification at any time
(PAD)	during study period (from baseline to follow-up).
	Glycated hemoglobin A1c \geq 6.5% or fasting glucose \geq 7.0 mmol/L (\geq 126
	mg/dL) or non-fasting glucose ≥ 11.1 mmol/L (≥ 200 mg/dL) or use of
Diabetes mellitus	glucose lowering drugs (ADA 2010 criteria). Self-report of physician
	diagnosed diabetes can be included.
	Identification at any time during study period (from baseline to follow-up).

Hypertension	Systolic blood pressure ≥140 mmHg or diastolic blood pressure ≥90 mmHg or antihypertensive drugs (JNC-7 criteria). Self-reported hypertension if other data not available. Identification at any time during study period (from baseline to follow-up).
History of cancer	Prior diagnosis of any cancer. Identification at any time during study period (from baseline to follow-up).
History of liver disease	Prior diagnosis of liver disease. Identification at any time during study period (from baseline to follow-up).
History of chronic obstructive pulmonary disease (COPD)	Prior diagnosis of COPD. Identification at any time during study period (from baseline to follow-up).

Outcome definitions below were provided to cohorts, cohort deviations from these definitions listed in the analytic notes for each cohort.

Outcome	Definition
All-cause mortality	Death
Cardiovascular mortality	Death due to CVD
Myocardial infarction	First incidence of myocardial infarction or fatal myocardial infarction
Stroke	First incidence of stroke (ischemic or hemorrhagic) or fatal stroke
Heart Failure	First hospitalization or death due to heart failure
Peripheral artery disease	First incidence of peripheral artery disease based on your study definition
Atrial fibrillation	First incidence of atrial fibrillation
Kidney failure with replacement	Initiation of dialysis or transplantation
therapy	
Acute kidney injury	First hospitalization of acute kidney injury
Hospitalizations	First hospitalization after baseline

ICD codes used to define outcomes within cohorts, if not specified in analytic notes

Outcome	ICD-9 codes	ICD-10 codes
Candiaya aayaa maatality	401-414, 426-443 (excluding 426.7,	I10-I25, I44-I73 (excluding I45.6, I51.4, I60,
Cardiovascular mortality	429.0, 430, 432.1, 437.3, 437.4, 437.5)	162.0, 167.1, 167.5, 167.7)
Myocardial infarction (MI)	410	I21, I22
Stroke	431, 432, 433.x1, 434.x1	I61, I62, I63
Heart failure	428	150
Dorinharal artary diagona	440.2, 440.3, 440.4, 38.18, 39.25,	170.2, 170.3, 170.4, 170.5, 170.92, 031, 041,
Peripheral artery disease	39.29, 39.50, 84.1	047, 04B, 04C, 04H, 04R, 04U, 0Y6
Atrial fibrillation	427.3	I48
Acute kidney injury	584	N17

The CKD-PC data request and processing procedures are as follows. After obtaining opt-in preferences from cohorts for the topics for each phase, the Data Coordinating Center (DCC) requests de-identified data using a specific data request document describing the variables and preferred definitions needed for the current phase of the CKD-PC. Cohorts work with the DCC on any data use agreements, IRB approvals, and other logistic issues for de-identified data transfer. The DCC also advises on any differences in definitions or questions on data formatting. Cohorts then provide de-identified data (in whatever program format, e.g., Stata, SAS, csv) via a secure data transfer provided or their own secure transfer program/platform. Data is stored on a secure password protected network server that is only accessed by limited faculty and staff (<10). All those faculty and staff have completed HIPAA and CITI certification and have signed internal data use agreements to not use the data for any other than stated purposed and to not remove the data from that network drive. The CKD-PC does not share data with any external parties. Once data is received and stored in the network drive, the DCC programmer reviews the data and the data dictionary provided by the cohort to check for any missing information, outliers, and potentials issues with variable units, dates, etc. Any questions are sent to the cohort representatives for data checking and cleaning. Further data checking is done throughout the analysis process for each CKD-PC paper, including a review from a cohort representative of all tables and figures to confirm their cohort representation.

For 98 of the 114 cohorts in this specific study, the DCC at Johns Hopkins University conducted the analysis; the remainder ran the standard code written in Stata by the DCC and shared the output with the DCC. As in the data processing procedures above, the DCC works with the cohort to confirm the variable definitions and data formatting to prepare for the code. The standard code was designed to automatically save all estimates and variance-covariance matrices needed for the meta-analysis. Then the DCC meta-analyzed the estimates across cohorts using Stata.

As detailed in our previous reports,^{2,3} each cohort was instructed to standardize their serum creatinine and report its method when available. The reported creatinine standardization allows grouping studies into studies that reported using a standard IDMS traceable method or conducted some serum creatinine standardization to IDMS traceable methods (ARIC, AusDiab, BIS, CanPREDDICT, CKD-JAC, CKD-REIN, COBRA, CRIC, ESTHER, GCKD, Geisinger, GLOMMS, Go-DARTS, Gubbio, ICES-KDT, ICKD, LCC, JHS, Maccabi, MASTERPLAN, MMKD, Nanjing CKD, NEPHROTEST, NHANES, Okinawa, PREVEND, PSPA, Rancho Bernardo, RCAV, REGARDS, SCREAM, SKS, SEED, SRR-CKD, STOP-CKDu, Takahata, West of Scotland, UK Biobank) and studies where the creatinine standardization was not done (AASK, ADVANCE, Aichi, CARE, BC CKD, Beijing, China NS, CHS, CIRCS, CRIB, Framingham, KP Hawaii, MDRD, MESA, MRC, NEFRONA, NIPPON DATA80/90, Ohasama, Pima, PSP-CKD, RENAAL, SMART, Sunnybrook, Taiwan MJ, TLGS, ULSAM, ZODIAC). For those cohorts without standardization, the creatinine levels were reduced by 5%, the calibration factor used to adjust non-standardized MDRD Study samples to IDMS.^{2,4}. We did not adjust creatinine levels in those studies with unknown standardization status (CURE-CKD, Gonryo, Hong Kong CKD, JMS, J-SHC, Mt Sinai BioMe, SHARP, NRHP-URU, YWSCC, and OLDW all cohorts).

Cohort	Notes including if a calibration equation was applied				
ARIC	IFCC Cystatin C = 1.12*(0.083+0.914*(ARIC cystatin C))				
AUSDIAB	IFCC Cystatin C = 1.12*(-0.25+1.07*(AusDiab cystatin C))				
BIS	ERM-DA471/IFCC Standardized assay				
CHS	IFCC= 1.12*(0.083+0.789*(CHS cystatin C))				
ESTHER	IFCC= 1.12*(0.105+0.848*(ESTHER cystatin C))				
FRAMINGHAM	IFCC= 1.12*(0.083+0.789*(Framingham cystatin C))				
MESA	IFCC= 1.12*(0.083+0.789*(MESA cystatin C))				
NHANES99-02	IFCC= 1.12*((NHANES1999-2002 cystatin C)-0.12)				
PREVEND	IFCC= 1.12*(0.083+0.789*(PREVEND cystatin C))				
REGARDS	Calibrated by primary study				
SCREAM	Calibrated by primary study				
UK BioBank	ERM-DA471/IFCC Standardized assay				
ULSAM	IFCC= 1.12*(0.083+0.789*(ULSAM cystatin C))				
AASK	IFCC= 1.12*(0.083+0.789*(AASK cystatin C))				
CKD-Rein	ERM-DA471/IFCC Standardized assay				
CRIB	IFCC= 1.12*(0.083+0.789*(CRIB cystatin C))				
CRIC	Calibrated by primary study				
GCKD	ERM-DA471/IFCC Standardized assay				
MASTERPLAN	IFCC=1.12*(MASTERPLAN cystatin C))				
MDRD	IFCC=1.12*(0.083+0.789*(MDRD cystatin C))				

Serum cystatin C values were calibrated and/or standardized to International Federation for Clinical Chemists
(IFCC) reference (Inker et al., 2011; Grubb et al., 2010). ^{5,6} Cohort details below:

		sing the 2021	CKD-EPI creatinine and the 2021 CKD-EPI creatinine-cystatin C equations, ⁷ as
follows:			
Sev	Serum	Serum	Equation

Sex	Serum	Serum	Equation
	Creatinine	Cystatin C	
	(mg/dL)	(mg/L)	
CKD-EF	PI creatinine of	equation	
Female	≤0.7		$GFR = 142 \text{ x} (Scr/0.7)^{-0.241} \text{ x} 0.9938^{Age}$
	>0.7		$GFR = 142 \text{ x} (Scr/0.7)^{-1.200} \text{ x} 0.9938^{Age}$
Male	≤0.9		$GFR = 142 \text{ x} (Scr/0.9)^{-0.302} \text{ x} 0.9938^{Age}$
	>0.9		$GFR = 142 \text{ x} (Scr/0.9)^{-1.200} \text{ x} 0.9938^{Age}$
CKD-EP	I creatinine-c	ystatin C equa	tion
Female	≤0.7	≤0.8	GFR= 135 x (Scr/0.7) ^{-0.219} x (Scys/0.8) ^{-0.323} x 0.9961 ^{Age} x 0.963
		>0.8	GFR= 135 x (Scr/0.7) ^{-0.544} x (Scys/0.8) ^{-0.778} x 0.9961 ^{Age} x 0.963
	>0.7	≤ 0.8	GFR= 135 x (Scr/0.7) ^{-0.219} x (Scys/0.8) ^{-0.323} x 0.9961 ^{Age} x 0.963
		>0.8	GFR= 135 x (Scr/0.7) ^{-0.544} x (Scys/0.8) ^{-0.778} x 0.9961 ^{Age} x 0.963
Male	≤0.9	≤0.8	$GFR = 135 \text{ x} (Scr/0.9)^{-0.144} \text{ x} (Scys/0.8)^{-0.323} \text{ x} 0.9961^{Age}$
		>0.8	GFR= 135 x (Scr/0.9) ^{-0.544} x (Scys/0.8) ^{-0.778} x 0.9961 ^{Age}
	>0.9	≤ 0.8	GFR= 135 x (Scr/0.9) ^{-0.144} x (Scys/0.8) ^{-0.323} x 0.9961^{Age}
		>0.8	GFR= 135 x (Scr/0.9) ^{-0.544} x (Scys/0.8) ^{-0.778} x 0.9961 ^{Age}

The selection of knots for eGFR and urine albumin-to-creatinine ratio was based on clinical thresholds.⁸ Baseline for each study was considered first available creatinine unless otherwise noted. Other variables were taken either on baseline date or within one year before baseline date.

Outcome	Model adjustment variables			
All-cause mortality	Age, sex, smoking status (current, former, never), systolic blood pressu body-mass index, use of anti-hypertensive medications, and a medical history of diabetes, coronary heart disease, stroke, heart failure, atrial fibrillation, peripheral artery disease, cancer, and chronic obstructive pulmonary disease			
Cardiovascular mortality	Age, sex, smoking status (current, former, never), systolic blood pressure, total cholesterol, high-density lipoprotein cholesterol, body-mass index, use of anti-hypertensive medications, and a medical history of diabetes, coronary heart disease, stroke, heart failure, atrial fibrillation, peripheral artery disease, cancer, and chronic obstructive pulmonary disease			
Myocardial infarction	Age, sex, smoking status (current, former, never), systolic blood pressure, total cholesterol, high-density lipoprotein cholesterol, body-mass index, use of anti-hypertensive medications, and a medical history of diabetes, stroke, heart failure, atrial fibrillation, peripheral artery disease, cancer, and chronic obstructive pulmonary disease			
Stroke	Age, sex, smoking status (current, former, never), systolic blood pressure, total cholesterol, high-density lipoprotein cholesterol, body-mass index, use of anti-hypertensive medications, and a medical history of diabetes, coronary heart disease, heart failure, atrial fibrillation, peripheral artery disease, cancer, and chronic obstructive pulmonary disease			
Heart Failure	Age, sex, smoking status (current, former, never), systolic blood pressure, total cholesterol, high-density lipoprotein cholesterol, body-mass index, use of anti-hypertensive medications, and a medical history of diabetes, coronary heart disease, stroke, atrial fibrillation, peripheral artery disease, cancer, and chronic obstructive pulmonary disease			

Each analysis was performed separately within each cohort. The models were adjusted for the following variables:

Peripheral artery disease	Age, sex, smoking status (current, former, never), systolic blood pressure,		
	total cholesterol, high-density lipoprotein cholesterol, body-mass index,		
	use of anti-hypertensive medications, and a medical history of diabetes,		
	coronary heart disease, stroke, heart failure, atrial fibrillation, cancer, and		
	chronic obstructive pulmonary disease		
Atrial fibrillation	Age, sex, smoking status (current, former, never), systolic blood pressure,		
	total cholesterol, high-density lipoprotein cholesterol, body-mass index,		
	use of anti-hypertensive medications, and a medical history of diabetes,		
	coronary heart disease, stroke, heart failure, peripheral artery disease,		
	cancer, and chronic obstructive pulmonary disease		
Kidney failure with replacement	Age, sex, smoking status (current, former, never), systolic blood pressure,		
therapy	body-mass index, use of anti-hypertensive medications, and a medical		
	history of diabetes, coronary heart disease, stroke, heart failure, atrial		
	fibrillation, peripheral artery disease, cancer, and chronic obstructive		
	pulmonary disease		
Acute kidney injury	Age, sex, smoking status (current, former, never), systolic blood pressure,		
	body-mass index, use of anti-hypertensive medications, and a medical		
	history of diabetes, coronary heart disease, stroke, heart failure, atrial		
	fibrillation, peripheral artery disease, cancer, and chronic obstructive		
	pulmonary disease		
Hospitalizations	Age, sex, smoking status (current, former, never), systolic blood pressure,		
respresizations	body-mass index, use of anti-hypertensive medications, and a medical		
	history of diabetes, coronary heart disease, stroke, heart failure, atrial		
	•		
	fibrillation, peripheral artery disease, cancer, and chronic obstructive		
	pulmonary disease		

Within each cohort, if a variable was missing more than 50% or not available, the variable was not included in the model. For variables that were missing <50% of the time, missing values were imputed with the mean.

By definition, age, sex, and eGFR were never missing.

Because albuminuria was a main exposure, different criteria were used. If albuminuria was missing <10%, complete case analysis was performed. Otherwise, missing albuminuria was analyzed as a separate "dummy" category. For estimation of hazard ratios in meta-analysis, the missing category was meta-analyzed only for EHR cohorts.

Details of missingness in each cohort are provided in the supplement Table S1.

1.2 Notes for individual cohorts:

Cohort	Study Design	Baseline Year(s)	Albuminuria type(s)*	Cystatin C available	Specific notes, including outcome ascertainment
General and High-Risk Coho	orts				
ADVANCE	Clinical trial cohort	2001-03	ACR	No	This study is a clinical trial which includes participants with diabetes only. All outcomes were actively ascertained and verified by an adjudication committee blinded to the randomized treatment.
Aichi	Worker health checkup database	2002-03	Dipstick	No	Reports from the worksite's healthcare division, and self- report with or without hospital records review were used to ascertain cardiovascular diseases incidence. The former and the reports of the next of kin were used to ascertain mortality.
ARIC	Research Cohort	1996-98	ACR	Yes	Visit 4 was used as the baseline. Kidney failure replacement therapy was ascertained through linkage to the USRDS. ⁹ All-cause mortality was actively ascertained as well as through linkage to a registry. Cardiovascular mortality was ascertained through review of ICD codes.
AusDiab	Research Cohort	1999-2000	ACR	Yes	Linkage to National Death Index for all-cause mortality and other fatal outcomes.
Beijing	Research Cohort	2004	ACR	No	All-cause mortality was actively ascertained.
BIS	Research Cohort	2009-11	ACR,	No	Kidney failure replacement therapy was ascertained by self- report during follow-up and validated by health records. All-cause mortality was actively ascertained through information of the insurance company and death certificates. Cardiovascular mortality was ascertained through death certificates. All other outcomes were self- reported.
CARE	Clinical trial cohort	1989-90	Dipstick	No	All outcomes were actively ascertained by an independent review committee.
China NS	Research cohort	2006-10	ACR	No	All-cause mortality was actively ascertained. Cardiovascular mortality was ascertained from medical chart review of death due to CHD or stroke.
CHS	Research Cohort	1996-97	ACR	Yes	Kidney failure replacement therapy was ascertained through linkage to a registry. All-cause mortality was actively ascertained. Cardiovascular mortality was

					ascertained through review of autopsies, death certificates, hospitalization records, and physician notes by the CHS Events Committee.
CIRCS	Research Cohort	1986-93	Dipstick	No	All-cause mortality was actively ascertained. Cardiovascular mortality was ascertained from medical chart review of fatal CHD or death within 30 days of an incident stroke. MI was ascertained by chart review. Stroke was ascertained by a physician panel reviewing medical charts and CT/MRI images for most cases.
COBRA	Research Cohort	2004-05	ACR	No	This study had no separate history of CHD, thus all participants with history of CVD were excluded from any of the analyses of CVD outcomes. Ascertainment of mortality and CVD mortality in COBRA was via active follow-up of participants and review of records (not linked registry).
ESTHER	Research Cohort	2000-02	Dipstick	Yes	All outcomes were actively ascertained by questionnaires sent to the study participants and their general practitioners 2, 5, 8, 11 and 14 years after the cohort's baseline assessment. In addition, fatal disease events were ascertained by a mortality register and ICD-10 codes of the leading cause of death on the death certificate.
Framingham	Research Cohort	1981-86	ACR	Yes	All outcomes from medical record review.
Geisinger	Healthcare administrative database	2008-19	ACR, PCR, Dipstick	No	Kidney failure replacement therapy was ascertained through linkage to a registry. No cause of death information available. Cardiovascular mortality was defined by any hospitalization with related ICD codes and died within 30 days. Baseline index date was set as the earliest date of a serum creatinine measurement after 2008 and at least one year after enrollment.
GLOMMS	Clinical database	2010-19	ACR, PCR	No	Kidney failure replacement therapy was ascertained through linkage to a registry. Baseline index date was set as the earliest date of a serum creatinine at least one year after enrollment.
Go-DARTs	Research cohort	2004-12	ACR	No	Kidney failure replacement therapy was ascertained through linkage to the Scottish Renal Registry plus eGFR<15 on two occasions at least 90 days apart. Baseline index date was set as the earliest date of a serum creatinine measurement at least one year after enrollment.

Gubbio	Research Cohort	1988-92	ACR	No	All outcomes were actively ascertained. Cardiovascular mortality was ascertained via ICD codes for MI based on hospital diagnoses or death certificates.
ICES-KDT	Healthcare administrative database	2008-17	ACR, PCR, Dipstick	No	Kidney failure replacement therapy was ascertained through linkage to a registry and ICD codes. Baseline index date was set as the earliest date of a serum creatinine measurement at least one year after enrollment.
JHS	Research Cohort	2000-04	ACR	No	All-cause mortality and cardiovascular mortality and events were ascertained through active follow-up in annual follow- up phone calls as well as review of medical records for hospitalizations. Cardiovascular events are validated and adjudicated.
JMS	Research Cohort	1992-95	Dipstick	No	
J-SHC	Health checkup database	2008	Dipstick	No	To ascertain all-cause mortality, in each district, used the birth date, sex, death date, and address code to identify the subject in both the National database of death certificates and National Health Insurance agency. If not clear in that first step, as the study does not have names, they asked for help from public health nurses in each district to confirm deaths. The study does not have the name in both databases, but the public health nurse has both information such as who participated and who died after the screening. Final data was verified by Chiho Iseki, Okinawa Heart and Renal Association. Cardiovascular mortality was ascertained using ICD-10 codes for cause of death.
KP Hawaii	Clinical database	2005-09	PCR	No	Kidney failure replacement therapy was actively ascertained.
Maccabi	Healthcare administrative database	2008-17	ACR	No	Urine albumin-to-creatinine ratio measures above 300 were imputed by PCR measures. First creatinine after 2008 was selected. Kidney failure replacement therapy was actively ascertained. No cause of death information available. Cardiovascular mortality was defined by any hospitalization with related ICD codes and died within 30 days. Baseline index date was set as the earliest date of a serum creatinine measurement after 2008 and at least one year after enrollment.
MESA	Research cohort	2000-02	ACR	Yes	All participants free from previous cardiovascular disease at baseline. There was no information of kidney failure replacement therapy in this study. All-cause mortality was

MRC	Research cohort	1995-99	Dipstick	No	 actively ascertained. Cardiovascular mortality was ascertained through review of hospital records or, for participants who experienced out-of-hospital cardiovascular deaths, review of interviews with next of kin and physicians by the MESA morbidity and mortality review committee. Outcomes were ascertained through linkage with national death and hospitalization records.
Mt Sinai BioMe	Clinical database	2008-15	ACR, PCR	No	Kidney failure replacement therapy was ascertained through ICD codes. Cardiovascular events were ascertained from ICD codes at hospital discharge. No mortality information available. Baseline index date was set as the earliest date of a serum creatinine measurement after 2008 and at least one year after enrollment.
NHANES	Research cohort	1988-2014	ACR	Yes	The information of kidney failure replacement therapy is not available in for this analysis. All-cause mortality was ascertained through linkage to National Death Index files. Cardiovascular mortality was ascertained by ICD codes.
NIPPON DATA80	Research cohort	1980	Dipstick	No	Cardiovascular mortality was ascertained by ICD codes from death certificates.
NIPPON DATA90	Research cohort	1990	Dipstick	No	Cardiovascular mortality was ascertained by ICD codes from death certificates.
Ohasama	Research cohort	1992-2011	Dipstick	No	All-cause mortality was actively ascertained. Cardiovascular mortality was ascertained from ICD codes for fatal CHD or fatal stroke. Stroke included fatal stroke from ICD codes and ischemic and hemorrhagic stroke adjudicated by a physician panel.
Okinawa	Health screening	1993-94	Dipstick	No	Kidney failure replacement therapy was ascertained through linkage to a registry.
OLDW	Healthcare administrative database	2012-2021	ACR, PCR, Dipstick	No	This study used de-identified electronic health record (EHR) data from the Optum Labs Data Warehouse (OLDW). The database contains longitudinal health information on enrollees and patients, representing a diverse mixture of ages, ethnicities and geographical regions across the United States. The EHR-derived data includes a subset of EHR data that has been normalized and standardized into a single database. ¹⁰ Cohort inclusion criteria was more than 50 events of any outcome before excluding missing values of main exposure variables. Smoking status might be under measured in this study. All outcomes were defined by ICD codes from encounters. No

Pima	Research	1982-2007	ACR	No	 cause of death information available. Cardiovascular mortality was defined by any hospitalization with related ICD codes and died within 30 days. Baseline index date was set as the earliest date of a serum creatinine measurement after 2008 and at least one year after enrollment. All-cause mortality was ascertained by linkage to the National Death Index. Cardiovascular mortality was adjudicated by review of all available clinical records and the ICD codes from the death certificate. Kidney failure
					replacement therapy was ascertained through review of clinical records.
PREVEND	Research cohort	1997-98	ACR	Yes	All-cause mortality was ascertained by linkage to a death registry. Cardiovascular mortality was ascertained by ICD codes from the death registry. Cardiovascular events were ascertained by ICD codes at hospital discharge.
Rancho Bernardo	Research cohort	1992-96	ACR	No	All-cause mortality was ascertained by linkage to a death registry. Cardiovascular mortality was ascertained by ICD codes from the death registry. Non-fatal cardiovascular events were ascertained by participant self-report.
RCAV	Healthcare administrative database	2004-2011	ACR, PCR	No	Kidney failure replacement therapy was ascertained through linkage to a registry. All other outcomes were defined by ICD codes from encounters. No cause of death information available. Cardiovascular mortality was defined by any hospitalization with related ICD codes and died within 30 days. Baseline index date was set as the earliest date of a serum creatinine at least one year after enrollment.
REGARDS	Research cohort	2003-07	ACR	Yes	Kidney failure replacement therapy was ascertained through linkage to USRDS. ⁹ All-cause mortality was actively ascertained. Cardiovascular mortality was ascertained through death certificates, medical records, and autopsy reports obtained to adjudicate cause of death.
SCREAM	Clinical database	2008-18	ACR, Dipstick	Yes	Kidney failure replacement therapy was actively ascertained through linkage to a registry that validates these endpoints through medical record comparison. All other non-mortality outcomes were defined by ICD codes from encounters. Baseline index date was set as the earliest date of a serum creatinine measurement after 2008 and at least one year after enrollment.

SEED	Research cohort	2004-11	ACR	No	All-cause mortality, cardiovascular mortality, acute myocardial infarction, stroke and kidney failure replacement therapy were ascertained by linkage to National Disease Registry and Death Registry.
SHARP	Clinical trial cohort	2003-2007	ACR	No	All outcomes were actively ascertained.
SMART	Research cohort	1996-2018	ACR	No	Kidney failure replacement therapy was actively ascertained (with chart validation). All-cause mortality was actively ascertained. Cardiovascular outcomes were adjudicated by a physician panel.
STOP-CKDu	Research cohort	2018	PCR	No	All-cause mortality was ascertained through active participant contact, medical record review, and verbal autopsy.
Taiwan MJ	Research cohort	1994-2011	Dipstick	No	Cardiovascular mortality was ascertained from ICD codes.
Takahata	Research cohort	2004	ACR, PCR, Dipstick	No	Cardiovascular mortality was ascertained from ICD codes from death certificates.
TLGS	Research cohort	1999-2001	Dipstick	No	Cardiovascular mortality was ascertained by cause of death adjudicated by a physician panel.
UK BioBank	Clinical database	2007-10	ACR	Yes	All-cause mortality was ascertained by linkage to a death registry. Cardiovascular mortality was ascertained by ICD codes from the death registry. All other outcome events were ascertained by ICD codes at hospital discharge.
ULSAM	Research cohort	1991-95	ACR	Yes	All-cause mortality was ascertained by linkage to a death registry. Cardiovascular mortality was ascertained by ICD codes from the death registry. All other outcome events were ascertained by ICD codes at hospital discharge.
ZODIAC	Research cohort	1998-2002	ACR	No	Mortality was ascertained by coupling information with the official death registration in the Netherlands, and where possible, ascertained cause of death with the GPs.
CKD Cohorts					
AASK	Clinical trial cohort	1995-98	PCR	Yes	This study only had black participants. Kidney failure replacement therapy and all-cause mortality were actively ascertained. There was no information of cardiovascular mortality.
BC CKD	Clinical database	2012	ACR, PCR	No	Kidney failure replacement therapy was actively ascertained.
CanPREDDICT	Research cohort	2008	ACR, PCR	No	Kidney failure replacement therapy was actively ascertained.

CKD-JAC	Research cohort	2007-08	ACR, PCR	No	All outcomes were actively ascertained. Cardiovascular events were adjudicated by an independent committee.
CKD-REIN	Research cohort	2013-15	ACR, PCR	Yes	All outcomes were actively ascertained. Furthermore, kidney failure with replacement therapy and all-cause mortality were also ascertained through linkage with national registries. Cardiovascular deaths were adjudicated by 2 cardiologists.
CRIB	Research cohort	1996-98	ACR, Dipstick	Yes	Kidney failure replacement therapy was actively ascertained (with chart validation).
CRIC	Research cohort	2003-08	ACR, PCR	Yes	Kidney failure replacement therapy was actively ascertained with chart validation as well as through linkage to a registry. All-cause mortality was actively ascertained as well as through linkage to a registry. Cardiovascular mortality was ascertained through adjudicated chart review.
CURE-CKD	Healthcare administrative database	2007-17	ACR, PCR	No	All outcomes were ascertained from ICD codes.
GCKD	Research Cohort	2010-12	ACR	Yes	All outcomes were actively ascertained (with confirmation in medical chart review).
Gonryo	Research Cohort	2006-08	PCR, Dipstick	No	All outcomes were actively ascertained.
Hong Kong CKD	Clinical database	2007-12	PCR	No	All outcomes were actively ascertained. Cardiovascular events and mortality were adjudicated by a physician panel.
ICKD	Research Cohort	2013-19	ACR	No	All outcomes were actively ascertained through direct contact with participants and medical record review.
LCC	Clinical database	2011	ACR, PCR	No	All-cause mortality was ascertained from medical records. All other outcomes ascertained through ICD codes from hospital discharge.
MASTERPLAN	Clinical trial cohort	2004-05	ACR, PCR	Yes	All outcomes were actively ascertained.
MDRD	Clinical trial cohort	1989-91	PCR	Yes	Kidney failure replacement therapy and all-cause mortality were actively ascertained as well as through linkage to a registry. Cardiovascular mortality was ascertained through review of ICD codes. Due to super low eGFR in this study, eGFR was modeled as linear term without knot.
MMKD	Research cohort	1997-98	PCR	No	Kidney failure replacement therapy was actively ascertained.
Nanjing CKD	Research cohort	2003-15	PCR, Dipstick	No	All outcomes were actively ascertained.

NEFRONA	Research cohort	2009-12	ACR, PCR	No	Participants free from previous cardiovascular disease at baseline. Cardiovascular events were ascertained by ICD codes from referring physicians.
NephroTest	Research cohort	2000-12	ACR, PCR	No	All-cause mortality and cardiovascular mortality and events were actively ascertained. Kidney failure replacement therapy was ascertained through linkage to a registry.
NRHP-URU	Clinical database	2001-14	ACR, PCR, Dipstick	No	Kidney failure replacement therapy was ascertained through linkage to a registry.
PSP-CKD	Clinical database	2010-13	ACR, PCR, Dipstick	No	All-cause mortality was ascertained from medical records. All other outcomes ascertained through ICD codes from hospital discharge.
PSPA	Research cohort	2009-10	PCR	No	Kidney failure replacement therapy was actively ascertained as well as through linkage to a registry. All- cause and cardiovascular mortality were actively ascertained.
RENAAL	Clinical trial cohort	1996-98	ACR	No	All outcomes were actively ascertained (with physician panel adjudication).
SKS	Research cohort	2002-16	PCR	No	All outcomes were actively ascertained (with physician panel adjudication).
SRR-CKD	Renal registry	2005-11	ACR	No	All-cause mortality was ascertained from medical records. Kidney failure replacement therapy was ascertained through linkage to a registry.
Sunnybrook	Clinical database	2001-10	ACR, PCR, Dipstick	No	Kidney failure replacement therapy was ascertained through linkage to a registry. All other outcomes ascertained through ICD codes from hospital discharge.
West of Scotland CKD	Clinical database	2009-18	ACR, PCR, Dipstick	No	Kidney failure replacement therapy, hospitalizations, myocardial infarction, stroke, heart failure, atrial fibrillation and peripheral arterial disease were actively ascertained and prospectively recorded by clinical staff in the renal electronic record. All-cause mortality was additionally ascertained by linkage to non-renal electronic health records.
YWSCC	Clinical database	2017	ACR, PCR, Dipstick	No	

*Type of albuminuria used in analyses with a preference for ACR. Does not necessarily indicate all types available within the cohort.

Appendix 2. Acronyms or abbreviations for cohorts included in the current study and their key references linked to the Web references

ADVANCE The Action in Diabetes and Vascular Disease: Preterax and Diamicron Modified Release Controlled Evaluation (ADVANCE) trial ¹¹ Aichi Aichi Workers' Cohort Study ¹² ARIC Atherosclerosis Risk in Communities Study ¹³ AusDiab Australian Diabetes, Obesity, and Lifestyle Study ¹⁴ Beijing Beijing Cohort Study ¹⁵ BIS Berlin Initiative Study ¹⁶ CARE The China National Survey of Chronic Kidney Disease CHIS Cardiovascular Health Study ¹⁶ CORA Control Of Blood Pressure and Risk Attenuation Study ²⁰ COBRA Control Of Blood Pressure and Risk Attenuation Study ²⁰ CORA Control Of Blood Pressure and Risk Attenuation Study ²⁰ CORA Control Of Blood Pressure and Risk Attenuation Study ²⁰ CORA Control Of Blood Pressure and Risk Attenuation Study ²⁰ CORA Control Of Blood Pressure and Risk Attenuation Study ²⁰ GENA Griediarge Teath Study ²² Geisinger Geisinger Health Study ²² Geisinger Geisinger Health Study ²² Geisinger Geisinger Health System ²³ GLOMMS Grampian Laboratory Outcomes, Morbidity and Mortality Studies ²⁴ Go-DARTs Genetics of Diabetes Audi	General population an	d High Cardiovascular Risk Cohorts
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West of Scotland CKD West of Scotland study ⁷⁸	SRR-CKD	Swedish Renal Registry CKD Cohort ⁷⁶
	Sunnybrook	Sunnybrook Cohort ⁷⁷
YWSCC Yonsei Wonju Severance CKD Cohort	West of Scotland CKD	West of Scotland study ⁷⁸
	YWSCC	Yonsei Wonju Severance CKD Cohort

Appendix 3. Acknowledgements and funding for collaborating cohorts

Cohort	List of sponsors
AASK	AASK was supported by grants to each clinical center and the coordinating center from the National Institute of Diabetes and Digestive and Kidney Diseases. In addition, AASK was supported by the Office of Research in Minority Health (now the National Center on Minority Health and Health Disparities, NCMHD) and the following institutional grants from the National Institutes of Health: M01 RR-00080, M01 RR-00071, M0100032, P20-RR11145, M01 RR00827, M01 RR00052, 2P20 RR11104, RR029887, and DK 2818-02. King Pharmaceuticals provided monetary support and antihypertensive medications to each clinical center. Pfizer Inc, AstraZeneca Pharmaceuticals, Glaxo Smith Kline, Forest Laboratories, Pharmacia and Upjohn also donated antihypertensive medications.
ADVANCE	ADVANCE was supported by research grants from Servier International and from the National Health and Medical Research Council (NHMRC) of Australia program grants 358395, 571281, 1052555 and 1149987 and project grant 211086
Aichi	KAKENHI (09470112, 13470087, 17390185, 18590594, 20590641, 20790438, 22390133, 26293153, 18H03057)
ARIC	The Atherosclerosis Risk in Communities study has been funded in whole or in part with Federal funds from the National Heart, Lung, and Blood Institute, National Institutes of Health, Department of Health and Human Services, under Contract nos. (HHSN268201700001I, HHSN268201700003I, HHSN268201700005I, HHSN268201700004I, HHSN2682017000021). The authors thank the staff and participants of the ARIC study for their important contributions.
AusDiab	We wish to thank the AusDiab Steering Committee for providing data from the AusDiab study. Funding from NHMRC of Australia, Grant #1007544.
BC CKD	BC Provincial Renal Agency, an Agency of the Provincial Health Services Authority in collaboration with University of British Columbia.
Beijing	The research for this study was supported by the Program for New Century Excellent Talents in University (BMU2009131) from the Ministry of Education of the People's Republic of China, and the grants for the Early Detection and Prevention of Non- communicable Chronic Diseases from the International Society of Nephrology Research Committee.
BIS	Foundation for Preventive Medicine of the KfH (Kuratorium für Heimdialyse und Nierentransplantation e.V. – Stiftung Präventivmedizin; <u>www.kfh-stiftung-</u> <u>praeventivmedizin.de</u>). Verband deutscher Nierenzentren (DDnÄ)
CanPREDDICT	N/A
CARE	Alberta Heritage Foundation for Medical Research/Alberta Innovates Health Solutions Interdisciplinary Team Grants Program
China NS	N/A
CHS	This research was supported by contracts HHSN268201200036C, HHSN268200800007C, HHSN268201800001C, N01HC55222, N01HC85079, N01HC85080, N01HC85081, N01HC85082, N01HC85083, N01HC85086, 75N92021D00006, and grants U01HL080295 and U01HL130114 from the National Heart, Lung, and Blood Institute (NHLBI), with additional contribution from the National Institute of Neurological Disorders and Stroke (NINDS). Additional support was provided by R01AG023629 from the National Institute on Aging (NIA). A full list of principal CHS investigators and institutions can be found at CHS-NHLBI.org.
CIRCS	N/A
CKD-JAC	The CKD-JAC Study was financially supported by Kyowa Kirin Co., Ltd.
CKD-REIN	CKD-REIN is funded by the Agence Nationale de la Recherche through the 2010 «Cohortes-Investissements d'Avenir » program (ANR-IA-COH-2012/3731) and by the 2010 national Programme Hospitalier de Recherche Clinique. CKD-REIN is also supported through a public-private partnership with Fresenius Medical Care and

	GlaxoSmithKline (GSK) since 2012 and Vifor France since 2018, Sanofi-Genzyme
	from 2012 to 2015, Baxter and Merck Sharp & Dohme-Chibret (MSD France) from 2012 to 2017, Amgen from 2012 to 2020, Lilly France from 2013 to 2018, Otsuka
	Pharmaceutical from 2015 to 2020, AstraZeneca from 2018 to 2021 and Boehringer
	Ingelheim France since 2022.
COBRA	Wellcome Trust, UK
CODIUI	British Renal Society Project Grant Award
CRIB	British Heart Foundation Project Grant Award.
CRIC	Bittish Heart Foundation Froger Grant Award.Funding for the CRIC Study was obtained under a cooperative agreement from National Institute of Diabetes and Digestive and Kidney Diseases (U01DK060990, U01DK060984, U01DK061022, U01DK061021, U01DK061028, U01DK060980, U01DK060963, U01DK060902 and U24DK060990). In addition, this work was supported in part by: the Perelman School of Medicine at the University of Pennsylvania Clinical and Translational Science Award NIH/NCATS UL1TR000003, Johns Hopkins University UL1 TR-000424, University of Maryland GCRC M01 RR- 16500, Clinical and Translational Science Collaborative of Cleveland, UL1TR000439 from the National Center for Advancing Translational Sciences (NCATS) component of the National Institutes of Health and NIH roadmap for Medical Research, Michigan Institute for Clinical and Health Research (MICHR) UL1TR000433, University of Illinois at Chicago CTSA UL1RR029879, Tulane COBRE for Clinical and Translational Research in Cardiometabolic Diseases P20 GM109036, Kaiser Permanente NIH/NCRR UCSF-CTSI UL1 RR-024131, Department of Internal Medicine, University of New Mexico School of Medicine Albuquerque, NM
CURE-CKD	R01DK119199.The CURE-CKD registry was supported by institutional funding from Providence St.Joseph health and the University Of California, Los Angeles and by grant 75D301-19-Q-69877 from the US Centers for Disease Control and Prevention.
ESTHER	Ministry of Research, Science and the Arts Baden-Württemberg (Stuttgart, Germany), Federal Ministry of Education and Research (Berlin, Germany), Federal Ministry of Family Affairs, Senior Citizens, Women and Youth (Berlin, Germany), Saarland state Ministry for Social Affairs, Health, Women and Family Affairs (Saarbrücken, Germany). Measurement of urinary albumin was funded by Dade- Behring, Marburg, Germany.
Framingham	NHLBI Framingham Heart Study (N01-HC-25195).
GCKD	The GCKD study is supported by grants from the Federal Ministry of Education and Research (Bundesministerium für Bildung und Forschung; www.bmbf.de), FKZ 01ER 0804, 01ER 0818, 01ER 0819, 01ER 0820, 01ER 0821, and 01ER 0822, and the Foundation for Preventive Medicine of the KfH (Kuratorium für Heimdialyse und Nierentransplantation e.V. – Stiftung Präventivmedizin; www.kfh-stiftung- praeventivmedizin.de) and corporate partners (for a list see www.gckd.org). The GCKD investigators gratefully acknowledge the expert support of all members of study staff, the dedicated contribution of all collaborating nephrologists (for a list of contributors and the 169 study sites, see www.gckd.org) and the support of patients participating in the study. The work of AK was supported by the Deutsche Forschungsgemeinschaft (DFG, German Research Foundation) Project ID 431984000 SFB 1453.
Geisinger	Geisinger Clinic; NIDDK R01DK100446
GLOMMS	GLOMMS was initially funded, in first version, by a grant from Chief Scientist Office CZH/4/656. GLOMMS was subsequently expanded with support from a starter grant from the Academy of Medical Sciences, Wellcome Trust; Medical Research Council, British Heart Foundation; Arthritis Research UK; the Royal College of Physicians; and Diabetes UK [SGL020\1076]; and a research training fellowship from the Wellcome Trust [102729/Z/13/Z]. The GLOMMS study also acknowledges support from the Grampian Data Safe Haven (DaSH) facility within the Aberdeen Centre for Health Data Science and the associated financial support of

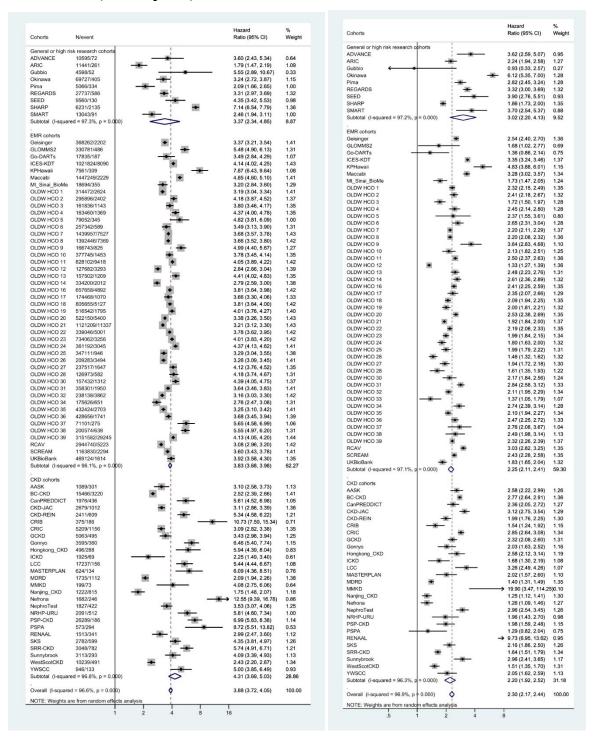
	the University of Aberdeen, and National Health Service (NHS) Research Scotland (through NHS Grampian investment in DaSH). More information is available at the
	DaSH website: http://www.abdn.ac.uk/iahs/facilities/grampian-data-safe-haven.php.
Go-DARTs	The Wellcome Trust United Kingdom Type 2 Diabetes Case Control Collection (supporting GoDARTS) was funded by the Wellcome Trust, under grants 072960/Z/03/Z, 084726/Z/08/Z, 084727/Z/08/Z, 085475/Z/08/Z, and 085475/B/08/Z.
Gonryo	This study was supported by grants from Astellas Pharm Inc. and the Miyagi Kidney Foundation.
Gubbio	Municipal and Health Authorities of Gubbio, Italy; Center of Gubbio Epidemiological Studies, Gubbio, Italy; University of Naples "Federico II", Naples, Italy.
Hong Kong CKD	This study was supported by the Hong Kong Health Service Research Funds and Fund support from Sanofi Renal.
ICES-KDT	This study was conducted at the ICES Western Site. ICES is funded by an annual grant from the Ontario Ministry of Health and Long-Term Care. ICES Western is funded by an operating grant from the Academic Medical Organization of Southwestern Ontario. This project was conducted with members of the provincial ICES Kidney, Dialysis and Transplantation Research Program (www.ices.on.ca/kdt), which receives programmatic grant funding from the Canadian Institutes of Health Research. Dr. Amit Garg is supported by the Dr. Adam Linton Chair in Kidney Health Analytics. This study was supported by ICES, which is funded by an annual grant from the Ontario Ministry of Health (MOH) and the Ministry of Long-Term Care (MLTC). The opinions, results and conclusions reported in this paper are those of the authors and are independent from the funding sources. No endorsement by ICES, the MOH or MLTC is intended or should be inferred. Parts of this material are based on data and/or information compiled and provided by CIHI and the Ontario Ministry of Health. The analyses, conclusions, opinions and statements expressed herein are solely those of the authors and do not reflect those of the funding or data sources; no endorsement is intended or should be inferred.
ICKD	This study is funded by a grant by the Department of Biotechnology, Government of India (No. BT/PR11105/MED/30/1345/2014).
JHS	The Jackson Heart Study (JHS) is supported and conducted in collaboration with Jackson State University (HHSN268201800013I), Tougaloo College (HHSN268201800014I), the Mississippi State Department of Health (HHSN268201800015I) and the University of Mississippi Medical Center (HHSN268201800010I, HHSN268201800011I and HHSN268201800012I) contracts from the National Heart, Lung, and Blood Institute (NHLBI) and the National Institute on Minority Health and Health Disparities (NIMHD). The authors also wish to thank the staffs and participants of the JHS.
JMS	N/A
J-SHC	This study was supported by a Health and Labor Sciences Research Grant for "Design of the Comprehensive Health Care System for Chronic Kidney Disease (CKD) based on the Individual Risk Assessment by Specific Health Checkup" from the Ministry of Health, Labor and Welfare of Japan and a Grant-in-Aid for "Research on Advanced Chronic Kidney Disease (REACH-J), Practical Research Project for Renal Disease" from the Japan Agency for Medical Research and Development (AMED).
KP Hawaii	N/A
LCC	Funded by the National Institute for Health Research (NIHR) Collaboration for Leadership in Applied Health Research and Care (CLAHRC) East Midlands and Kidney Research UK (Grant TF2/2015)
Maccabi	N/A
MASTERPLAN	The MASTERPLAN study is a clinical trial with trial registration ISRCTN registry: 73187232. Sources of funding: The MASTERPLAN Study was supported by grants from the Dutch Kidney Foundation (Nierstichting Nederland, number PV 01), and the

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MESA	This research was supported by contracts HHSN268201500003I, N01-HC-95159, N01-HC-95160, N01-HC-95161, N01-HC-95162, N01-HC-95163, N01-HC-95164, N01-HC-95165, N01-HC-95166, N01-HC-95167, N01-HC-95168 and N01-HC-95169 from the National Heart, Lung, and Blood Institute and by grants UL1-TR-000040 and UL1-TR-001079 from NCRR. The authors thank the other investigators, the staff, and the participants of the MESA study for their valuable contributions. A full list of participating MESA investigators and institutions can be found at http://www.mesa-nhlbi.org.
MMKD	The MMKD study was funded by the Austrian Heart Fund and by the Innsbruck Medical University.
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Supplemental Figure 1. Forest plot of hazard ratios associated with kidney failure with replacement therapy, stratified into general population cohorts, electronic health record cohorts, and CKD cohorts: from the continuous model: eGFR (first panel) and albuminuria (second panel)



*The forest plot depicts adjusted hazard ratios derived in each cohort using the continuous model depicted in Table 3. The first panel depicts the coefficient for the spline term for eGFR <60 ml/min/1,73 m2, expressed per 15

ml/min/1.73 m2 lower eGFR value. The second panel reflects the coefficient for the albuminuria coefficient, expressed per 8-fold higher albuminuria.

**N/n also included people missing albuminuria, which was modeled with an indicator variable. The median cohort % with albuminuria was 66 (6.3-100%) in the analytical eGFRcr population.

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