

# Chronic Kidney Disease – Detect, Protect, Perfect

7<sup>th</sup> March 2024 12.00-13:00



# House Keeping

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# The Health Innovation Network

The Academic Health Science Network for the North East and North Cumbria has changed its name to Health Innovation North East and North Cumbria (HI NENC).

The new name – which came into effect on 1st October following the start of our new five-year licence – reflects the organisation's key role to continue to support the development and spread of innovation across the region's health service.

But while our name has changed, our vision remains the same: to improve health outcomes, reduce inequalities, and boost the regional economy. Working alongside partners across the system, we will continue to accelerate health innovation in the region, and beyond.

Established in 2013 by NHS England we are one of 15 Health Innovations.



# Agenda

12.00 – 12.10 facts.ckd and HIN Strategic Priorities NENC  
Prof Julia Newton (Chair)

12.10 – 12.40 The link between CKD and CVD (including optimised management of long-term conditions e.g. hypertension, ASCVD, diabetes and heart failure).  
Prof Raj Thakkar

12.40 – 12.50 – KFRE and When to refer to Renal Service  
Dr Sarah McCloskey

12.50 – 13.00 – Q&A

13.00 - Close



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# Addressing key risk factors for cardiovascular disease: **chronic kidney disease**

[pccsuk.org](http://pccsuk.org)

## **Prof. Raj Thakkar**

GP  
President and CKD lead, Primary Care Cardiovascular Society  
Honorary Visiting Professor, Cardiff University Medical School  
Primary care cardiology lead, Oxford and Thames Valley Health Innovation Network  
Observing board member, British Society of Heart Failure  
Member, National Expert advisory groups for lipids, heart attack and HF/HVD, NHSE  
National primary care workstream co-lead - cardiac transformation programme, NHSE  
UK Clinical Director, Healthy.io  
Industry consultant

# Declarations

- Abbott
- AstraZeneca
- Bayer
- Boehringer Ingelheim
- Daiichi-Sankyo
- Novartis
- Amgen
- Medtronic
- Omron



**The PCCS are providers of education based on current guidance. We bear no responsibility for any adverse outcomes to patients which may occur with use of any therapies discussed during this presentation.**



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# Key topics

CKD as a risk factor for CVD

The importance of ACR testing [Prev/Sev/Risk]

How to improve outcomes



# What is Chronic Kidney Disease?

“The presence of kidney damage,  
mainly albuminuria

and/or

decreased kidney function  
(estimated glomerular filtration  
rate [eGFR]  $<60$  mL/min/1.73m<sup>2</sup>)

for at least 3 months”





# CKD Epidemic:



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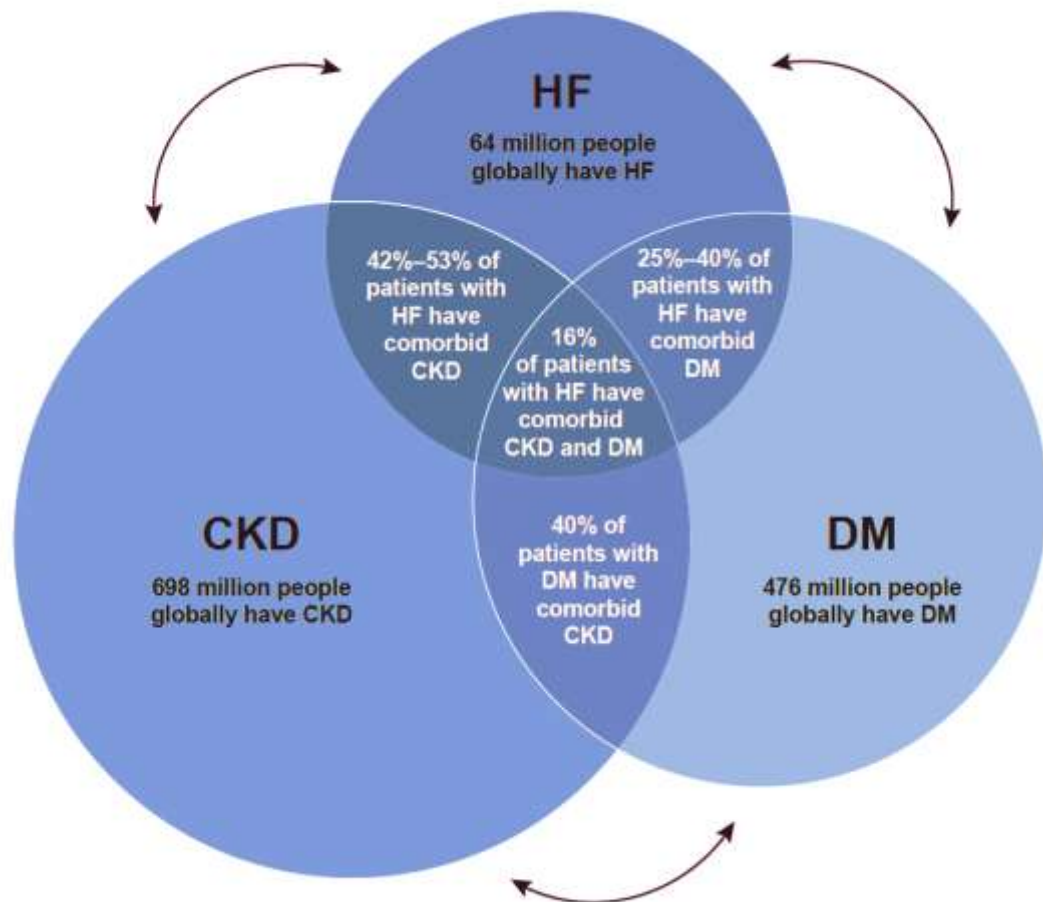
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- Global prevalence of CKD has risen by 87% between 1990–2016
- Estimated further increase of 17% in prevalence of CKD by 2030
- 3<sup>rd</sup> fastest growing cause of death
- 5<sup>th</sup> ranked cause of death by 2040
- UK: 2020 (3.63 million) 2030 (4.38 million) (Xie et al., 2018)
- 34% of CKD cases are undiagnosed
- Higher rates of CKD in under-served areas
- South Asians with diabetes 10x more likely to get kidney failure than Caucasians with diabetes



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## Heart Failure in Patients with Diabetes and Chronic Kidney Disease: Challenges and Opportunities

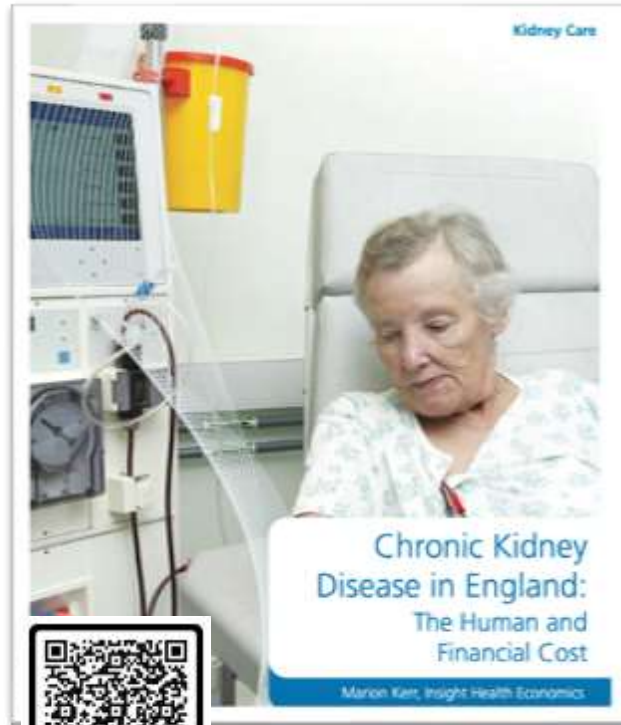


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### Review Article



## Health Economics of CKD:



SCAN ME

- **NHS England spent an estimated £1.45 billion on CKD in 2009–10**: equivalent to £1 in every £77 of NHS expenditure. This spending estimate covers both treatment directly associated with CKD (renal care and prescribing to prevent disease progression), and also treatment for excess non-renal problems such as strokes, heart attacks and infections in people with CKD.
- **There were an estimated 7,000 extra strokes and 12,000 extra myocardial infarctions in people with CKD in 2009–2010**, relative to the expected number in people of the same age and sex without CKD. The cost to the NHS of health care related to these strokes and MIs is estimated at £174–178 million.
- **People with CKD have longer hospital stays** than people of the same age without the condition, even when they go into hospital for treatments unrelated to CKD. We estimate that the average length of stay is 35% longer for people with CKD, and that the cost to the NHS of excess hospital bed days for patients with CKD was £46 million in 2009–10.



# Diagnosing and Classifying CKD [NICE, CKD 2021]:

Requires **both** blood testing [eGFR] and urine testing [ACR] to investigate patients for CKD

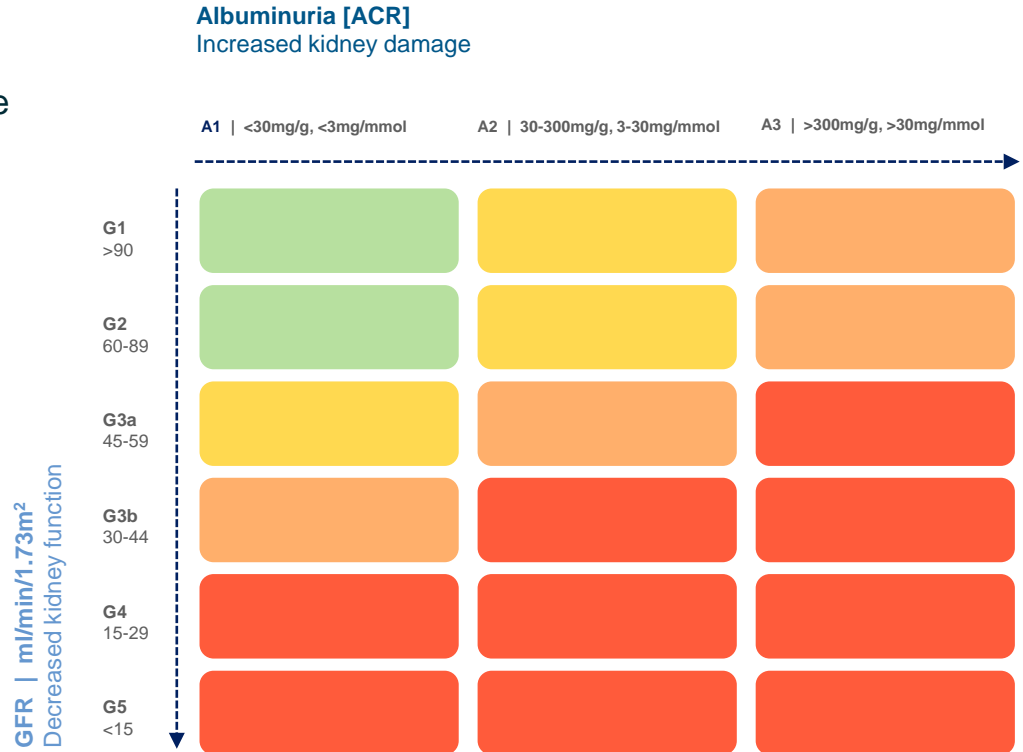
Key:

Low risk

Medium risk

High risk

Very high risk



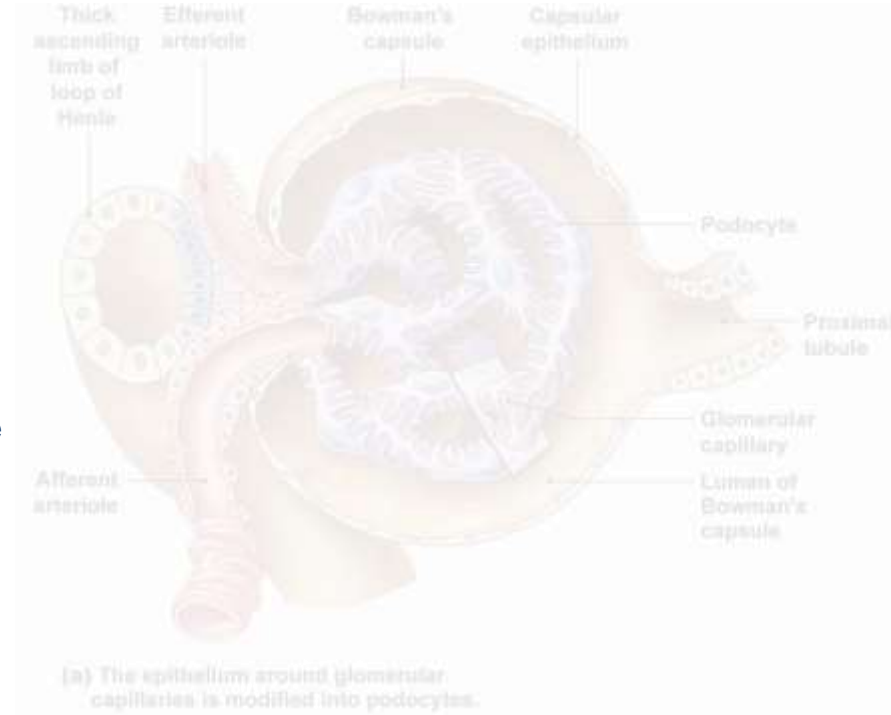


## Who should be tested for CKD?

### NICE NG203, CKD

1.1.21: Offer testing for CKD using eGFR and ACR to adults with any of the following risk factors:

- diabetes
- hypertension
- previous episode of acute kidney injury
- cardiovascular disease
- structural renal tract disease inc. stones, prostate disease
- gout
- multisystem diseases – e.g. SLE
- family history of end-stage renal disease (GFR category G5) or hereditary kidney disease



# The association of CKD with Cardiovascular Disease

The background features a dark blue gradient with several overlapping circles of varying shades of blue and grey. A small, light blue horizontal line is positioned to the left of the main text.



# For every 100 patients with moderate to severe Chronic Kidney Disease:

38 

unplanned hospital  
admissions per year

7 

events of acute  
kidney injury per year

2 

admissions to the  
Intensive Care Unit  
per year

6 

cardiovascular  
events per year

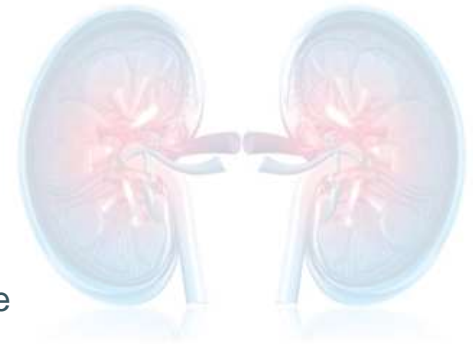
7 

deaths per year



# Kidney vasculature is a lens into the body's cardiovascular health:

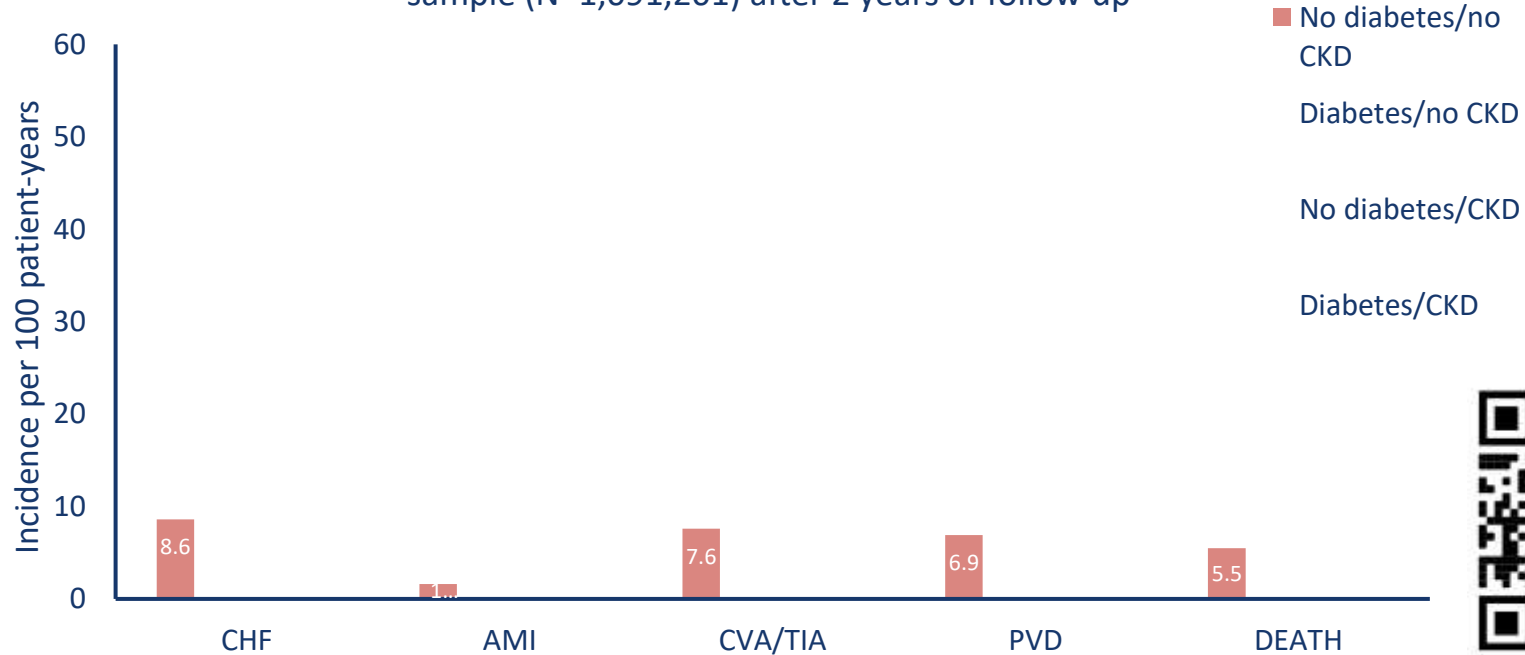
- Approximately 10 km of capillaries in both kidneys
- 180L plasma filtered by kidneys in 24 hours
- 20-25% cardiac output
- CKD is a cardiovascular risk state
- Patients with CKD are 20x more likely to die from CVD than renal failure
- **CKD must be considered one of the strongest risk factors for the development of CVD\***





# CKD vs diabetes

Incident event rates in 1998–1999 US Medicare population sample (N=1,091,201) after 2 years of follow-up

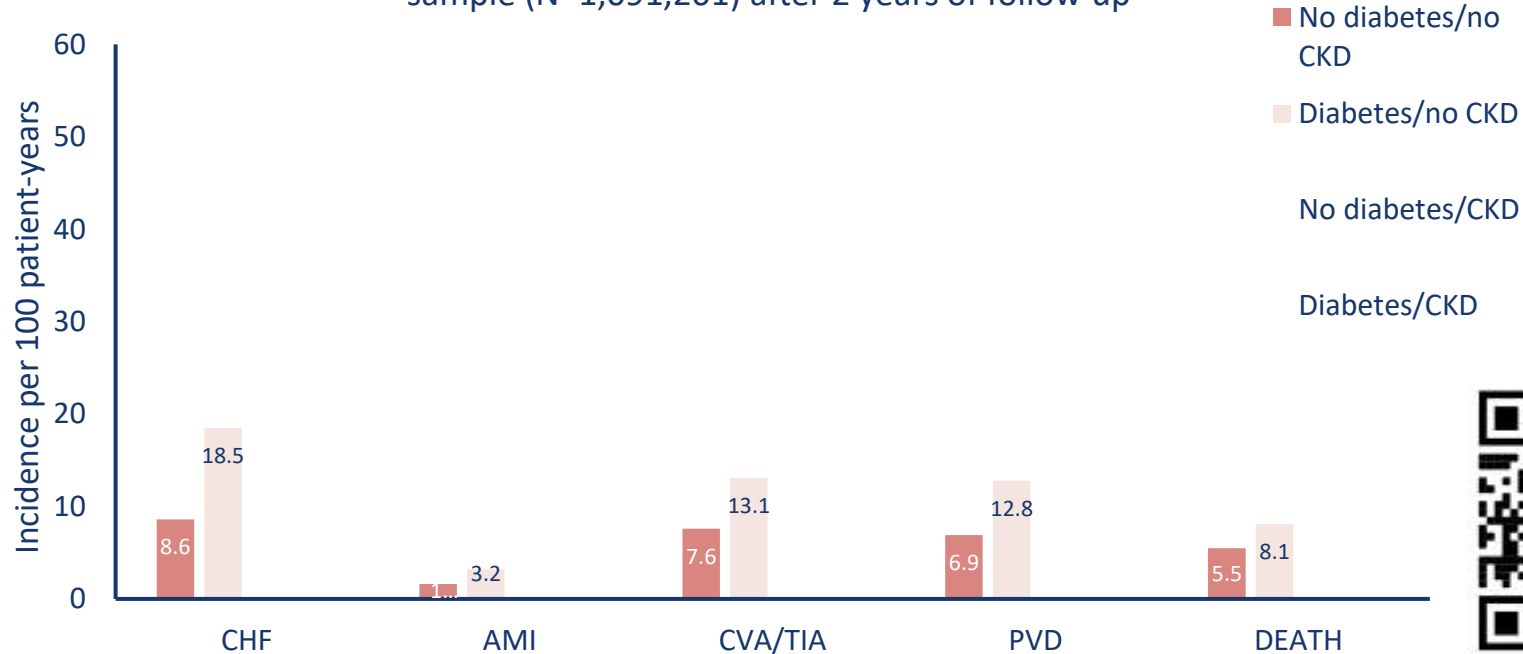


Increases in the values for Diabetes/no CKD to the values for Diabetes/CKD are indicated on the graph.



# CKD vs diabetes

Incident event rates in 1998–1999 US Medicare population sample (N=1,091,201) after 2 years of follow-up

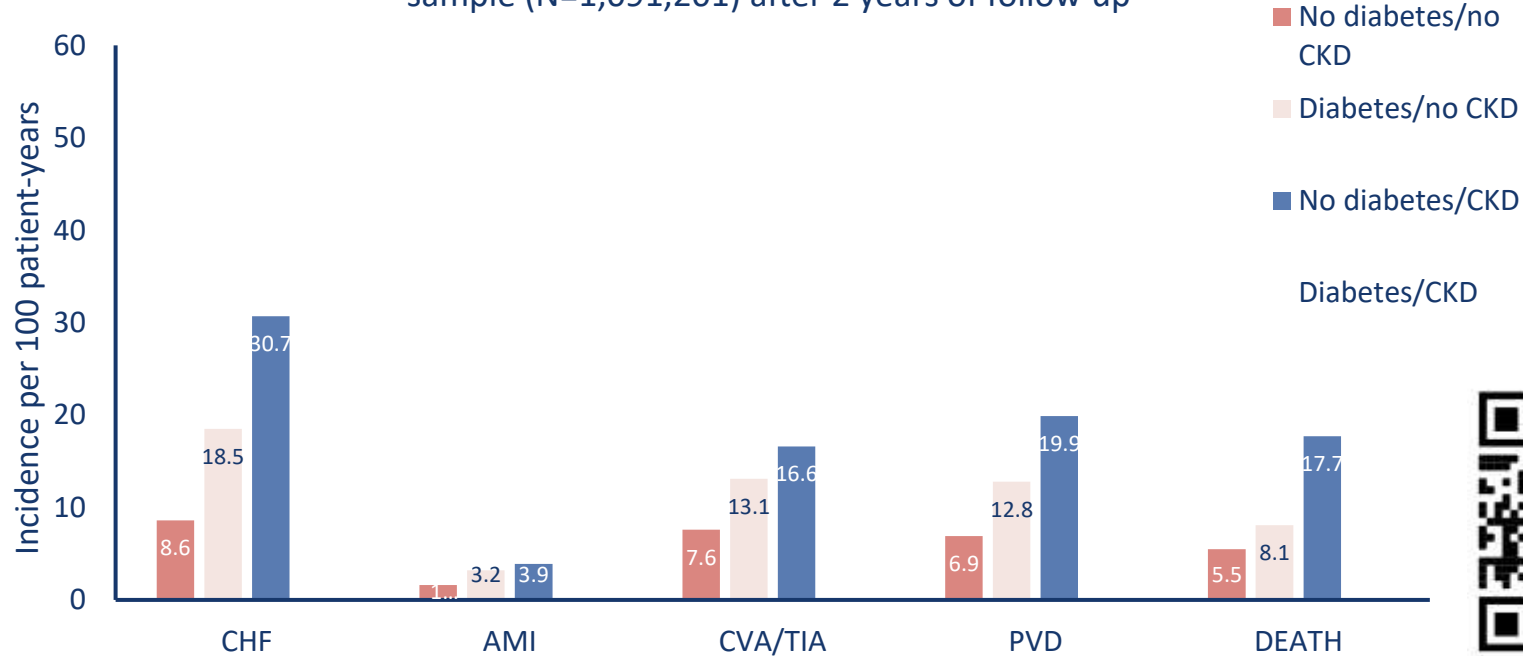


Increases in the values for Diabetes/no CKD to the values for Diabetes/CKD are indicated on the graph.



# CKD vs diabetes

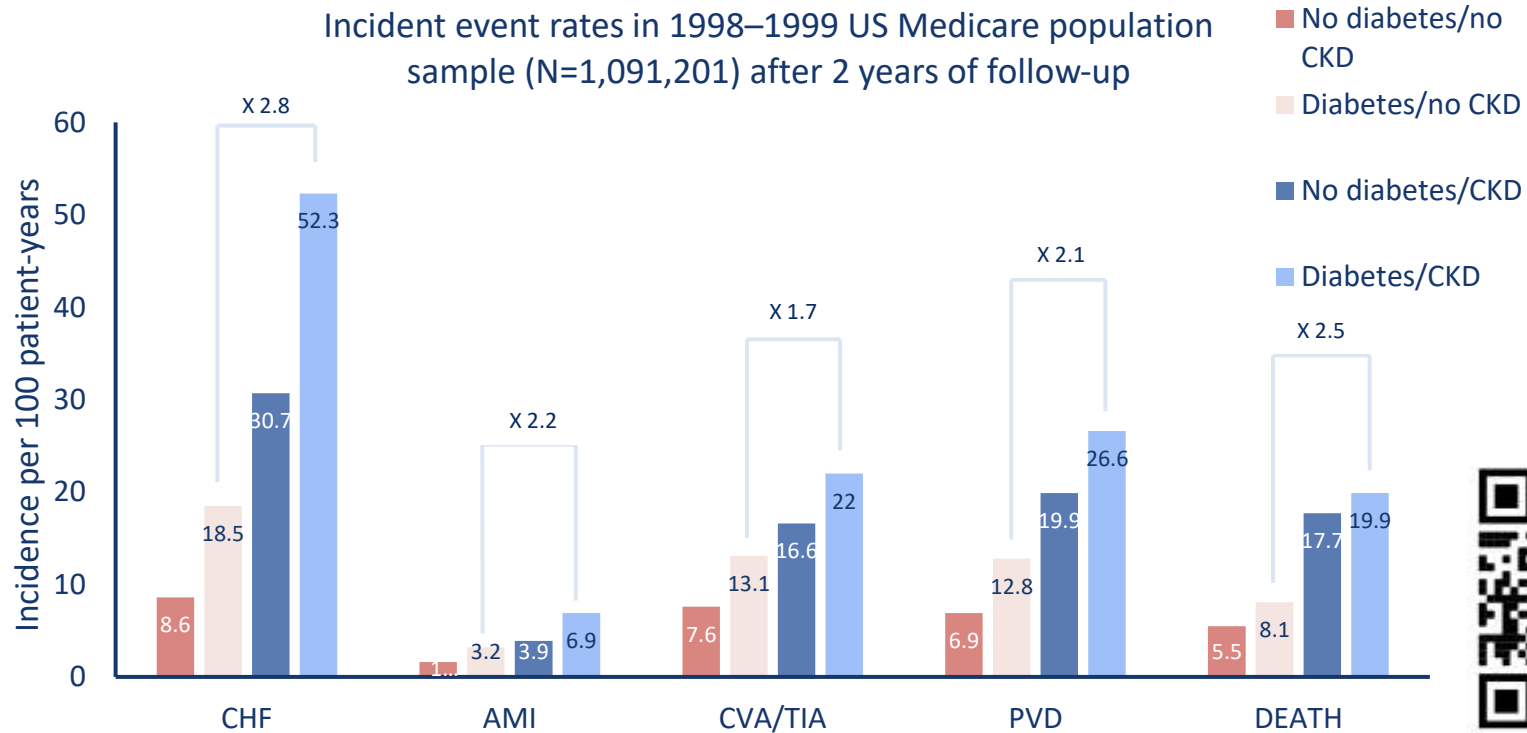
Incident event rates in 1998–1999 US Medicare population sample (N=1,091,201) after 2 years of follow-up



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# CKD vs diabetes

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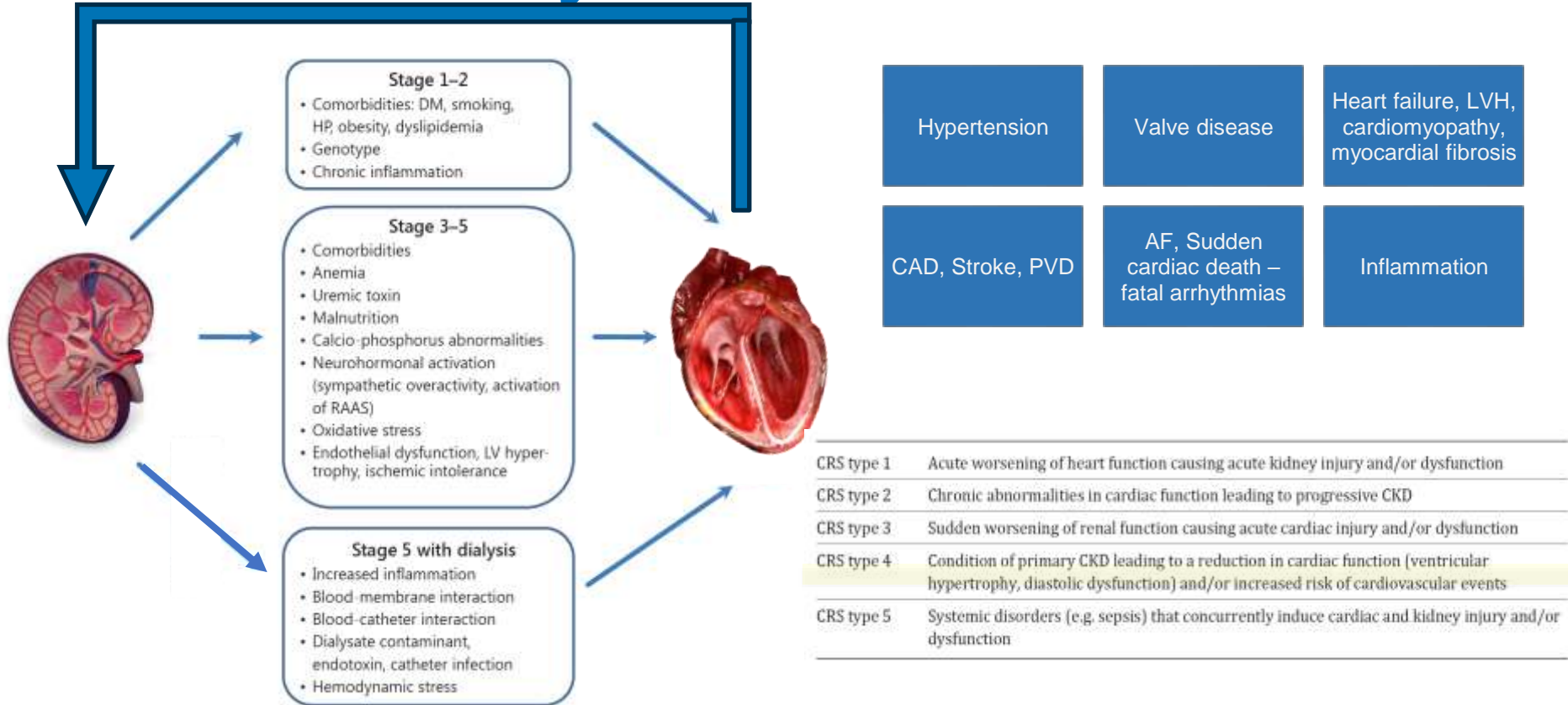


# Diabetic kidney disease

- ~40% diabetics develop DKD<sup>1-3</sup>
  - DKD is the commonest cause of ESRD worldwide
  - DKD in T2DM often diagnosed late
  - Often co-exists with obesity and AF
- – Significant number have albuminuria.<sup>4,5</sup>
  - Worse with uncontrolled HTN
  - Improves with good sugar and BP control
- – Complex<sup>2</sup>
  - Inflammation
  - RAAS activation and glomerular hyperfiltration
  - Glomerulosclerosis, fibrosis

1. Hussain S et al. Clinical Epidemiology and Global Health 2021;9:2–6; 2. Alicic RZ et al. Clin J Am Soc Nephrol. 2017;12:2032–2045; 3. Seyed Ahmadi S et al. Cardiovasc Diabetol 2020;19:9; 4. Selby NM et al. Diabetes Obes Metab. 2020;22 Suppl 1:3–15; 5. Thomas MC et al. Nat Rev Dis Primers. 2015;1:15018; 6. Wang et al., Atrial Fibrillation and Diabetes Mellitus: JACC Review Topic of the Week. Journal of the American College of Cardiology 2019;74:1107-1115.

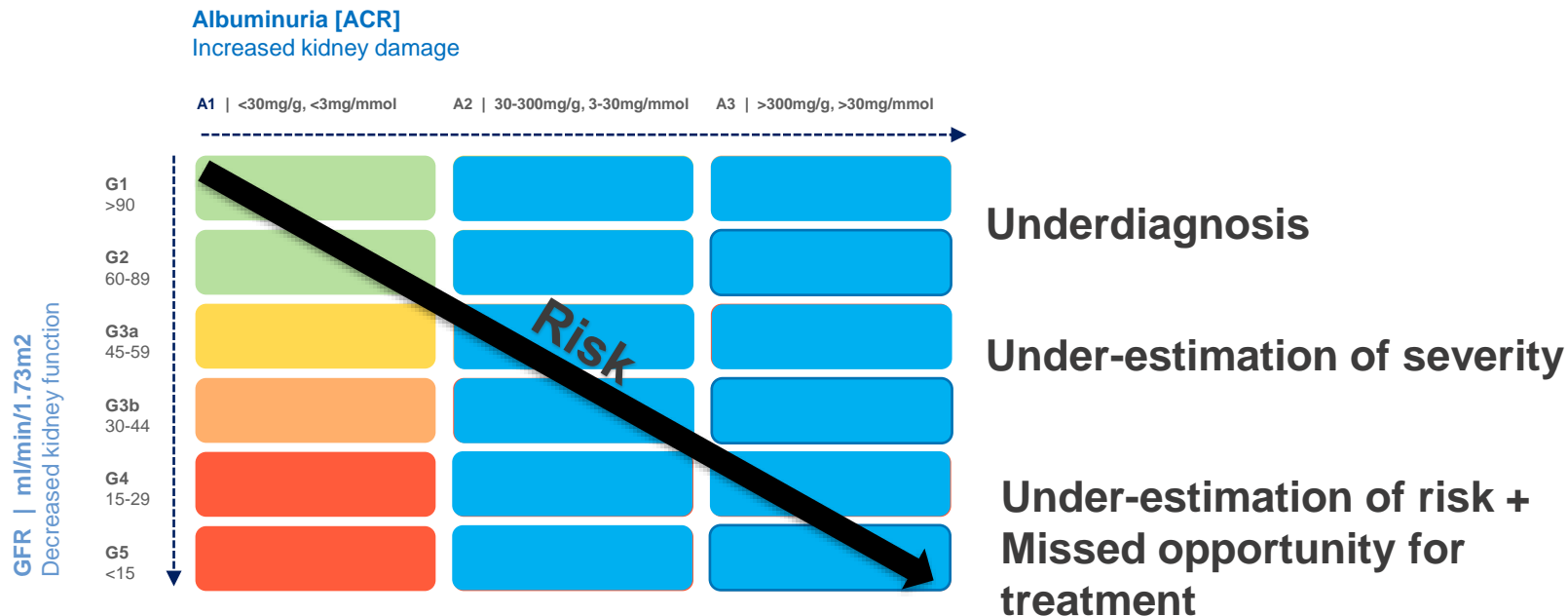
# Cardiorenal syndromes



AF, atrial fibrillation; CAD, coronary artery disease; CKD, chronic kidney disease; CRS, cardiorenal syndrome; DM, diabetes mellitus; HP, hypertension; LV, left ventricular; RAAS, renin-angiotensin-aldosterone system



# What happens if we don't check the urine for albuminuria?



Key

Low risk

Medium risk

High risk

Very high risk

# Albuminuria is a strong and independent risk predictor for end-stage renal disease (ESRD), CVD and death

Adjusted relative risk (RR) for eGFR cohorts by uACR level

All-cause mortality

	uACR < 1.0	uACR 1.0–2.9	uACR 3.0–29.9	uACR ≥ 30.0
eGFR > 105	1.1	1.5	2.2	5.0
eGFR 90–105	Ref	1.4	1.5	3.1
eGFR 75–90	1.0	1.3	1.7	2.3
eGFR 60–75	1.0	1.4	1.8	2.7
eGFR 45–60	1.3	1.7	2.2	3.6
eGFR 30–45	1.9	2.3	3.3	4.9
eGFR 15–30	5.3	3.6	4.7	6.6

Cardiovascular mortality

	uACR < 1.0	uACR 1.0–2.9	uACR 3.0–29.9	uACR ≥ 30.0
eGFR > 105	0.9	1.3	2.3	2.1
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eGFR 30–45	2.2	2.7	3.4	5.2
eGFR 15–30	14	7.9	4.8	8.1

Renal failure (ESRD)

	uACR < 1.0	uACR 1.0–2.9	uACR 3.0–29.9	uACR ≥ 30.0
eGFR > 105	Ref	Ref	7.8	18
eGFR 90–105	Ref	Ref	11	20
eGFR 75–90	Ref	Ref	3.8	48
eGFR 60–75	Ref	Ref	7.4	67
eGFR 45–60	5.2	22	40	147
eGFR 30–45	56	74	294	763
eGFR 15–30	433	1044	1056	2286

CVD, cardiovascular disease; eGFR, estimated glomerular filtration rate; ESRD, end-stage renal disease; uACR, urine albumin-to-creatinine ratio.

Adapted from Levey AS, et al. The definition, classification, and prognosis of chronic kidney disease: a KDIGO Controversies Conference report. *Kidney Int.* 2011; 80:17–28.



# Identification and management in primary care

## Identification

- CKD coding
- Case finding for unidentified CKD using eGFR and ACR
- Inequalities

## Management

- Education – Cardiovascular health / lifestyle / modifiable risk-factors
- QRISK3
- KFRE

- Lipid lowering therapy [QOF]
- Maximum Renin Angiotensinogen Aldosterone inhibition [Stop-ACEi study]
- Sodium Glucose Transporter-2 inhibitor and finerenone
- Optimise LTC
- Frailty/EOL

## Medical Optimisation

- Blood Pressure Optimisation



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# The problem with QRISK3



**About you**

Age (25-84):

Sex:  Male  Female

Ethnicity:

UK postcode: leave blank if unknown  
Postcode:

**Clinical information**

Smoking status:

Diabetes status:

Angina or heart attack in a 1st degree relative < 60?

Chronic kidney disease (stage 3, 4 or 5)?

Atrial fibrillation?

On blood pressure treatment?

Do you have migraines?

Rheumatoid arthritis?

Systemic lupus erythematosus (SLE)?

Severe mental illness?  
(this includes schizophrenia, bipolar disorder and moderate/severe depression)

On atypical antipsychotic medication?

Are you on regular steroid tablets?

A diagnosis of or treatment for erectile dysfunction?

Leave blank if unknown

Cholesterol/HDL ratio:

Systolic blood pressure (mmHg):

Standard deviation of at least two most recent systolic blood pressure readings (mmHg):

Body mass index

Height (cm):

Weight (kg):

**Welcome to the QRISK<sup>®</sup>3 risk calculator**

This site calculates a person's risk of developing a heart attack or stroke over the next 10 years, producing the score described in this academic paper:

- [Development and validation of QRISK3 risk prediction algorithms to estimate future risk of cardiovascular disease: prospective cohort study. BMJ 2017;357:j2099](#)

It presents the average risk of people with the same risk factors as those entered for that person.

The algorithm has been developed by doctors and academics working in the UK National Health Service and is based on routinely collected data from many thousands of GPs across the country who have freely contributed data to the QRResearch database for medical research.

It has been developed for the UK population, and is intended for use in the UK. All medical decisions need to be taken by a patient in consultation with their doctor. The authors and the sponsors accept no responsibility for clinical use or misuse of this score.

**Has QRISK<sup>®</sup>3 been validated?**

Yes. Validation of the underlying algorithm is described in the academic paper linked above. The software used to create this site has been tested using millions of randomly generated patient data (that is, simulated, not real data). Scores on this data match those generated by the statistical software used in the validation of the algorithm described in the academic paper.

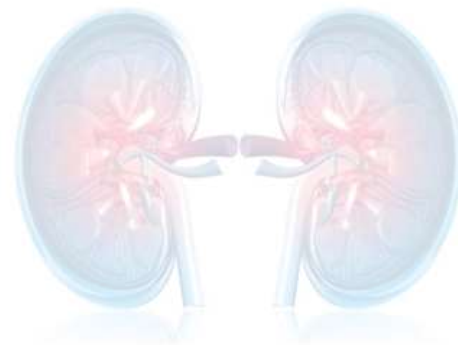
**Cardiovascular mortality**

	uACR < 1.0	uACR 1.0-2.9	uACR 3.0-29.9	uACR ≥ 30.0
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# Kidney failure risk equation

- Kidney failure risk equation = KFRE
- Adopted for UK population. - [www.kidneyfailurerisk.co.uk/](http://www.kidneyfailurerisk.co.uk/)
- Gives 5-year risk of end stage renal failure
- 5% referral threshold
- Doesn't give CVD risk

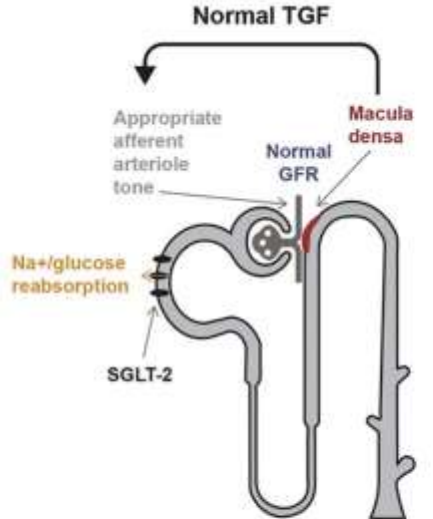


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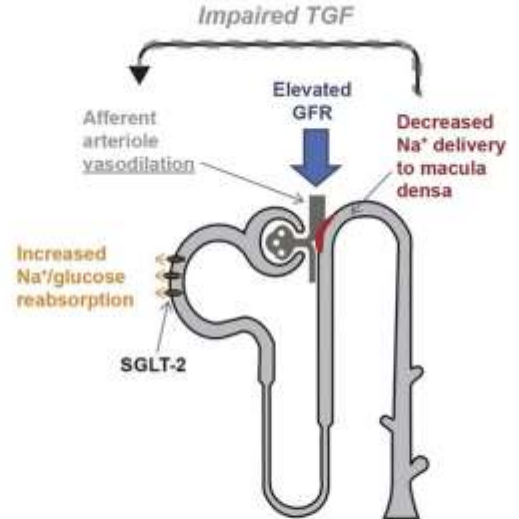
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# SGLT2i in diabetes

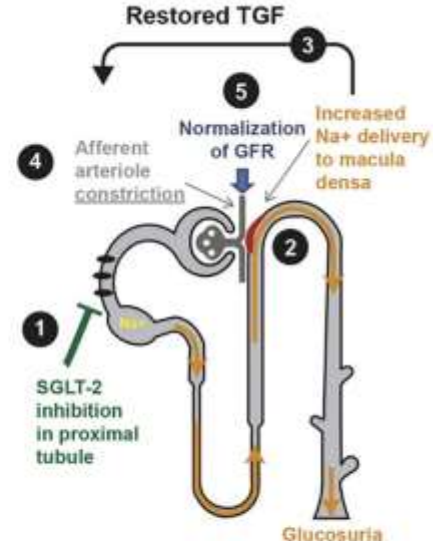
TGF: tubule glomerular feedback



Normal physiology



Hyperfiltration in early stages of diabetic nephropathy



SGLT-2 inhibition reduces hyperfiltration via TGF



# Quality improvement ideas in CKD

## Population health



- Code patients with CKD
- Actively look for patients at risk of having CKD using eGFR and ACR
- Ensure patients with CKD are auscultated for valve disease
- Have a high index of suspicion for heart failure

## Management



- Ensure patients with CKD esp with albuminuria are optimised
- Optimise CVD risk at an early stage
- Optimise secondary prevention
- Identify frailty early

# Summary

CKD is a strong predictor of adverse cardiovascular outcomes

CKD is a greater risk factor for CVD than diabetes

It is important to look for CKD in at risk patients [NICE].

Testing for eGFR alone is not enough –failure to test for albuminuria underestimates prevalence, severity of CKD and risk

Coding patients with CKD can reduce admissions and death

Address underlying cause, lifestyle factors, ACEi/ARBs, SGLT2i, finerenone, LLT and BP control

Identify frailty early

Thank you for your attention

 @DrRajThakkar



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**South Tyneside and Sunderland**  
NHS Foundation Trust

## CKD Webinar

Dr Sarah McCloskey  
Consultant Nephrologist and Physician  
NENC Renal Network CKD Lead

The path to  
**excellence**

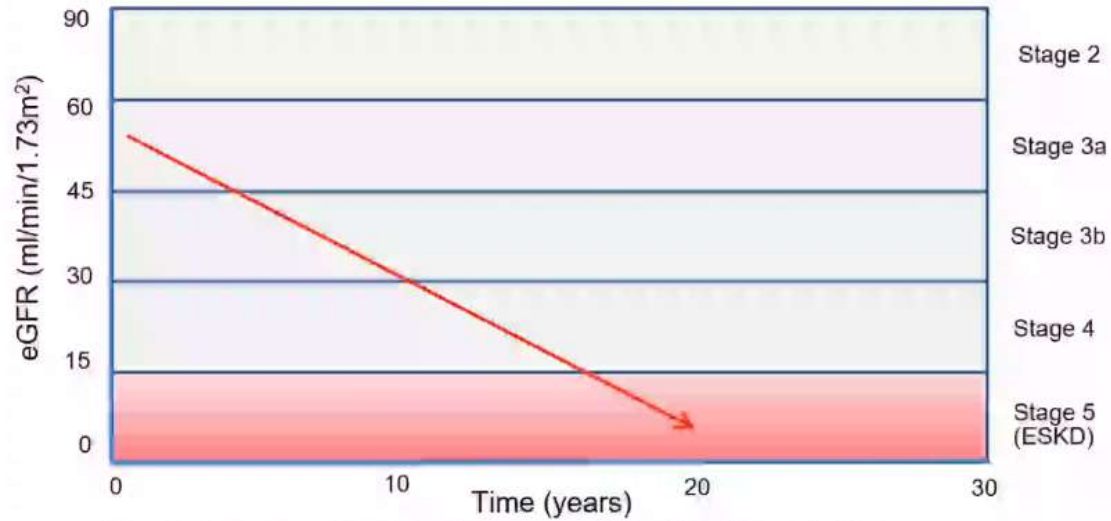


# Why does CKD matter?

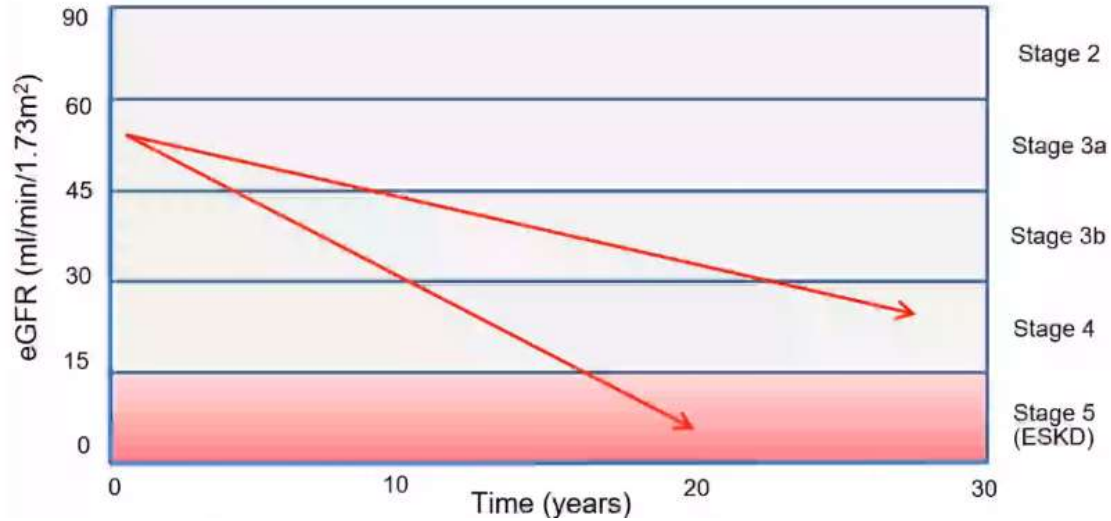
- CKD is a lifetime illness
- ↑ rate of mental health illnesses
- Lower QoL across all domains (esp. dialysis)
- ↓ employment prospects
- Impact on fertility in ESRD

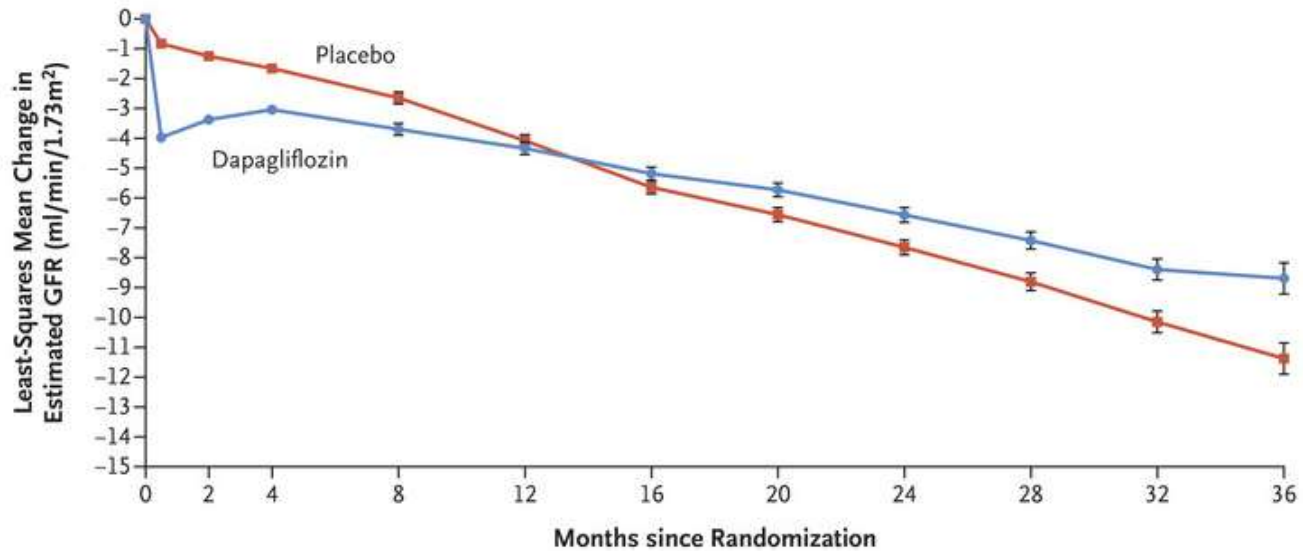


# Early identification and intervention



# Early identification and intervention

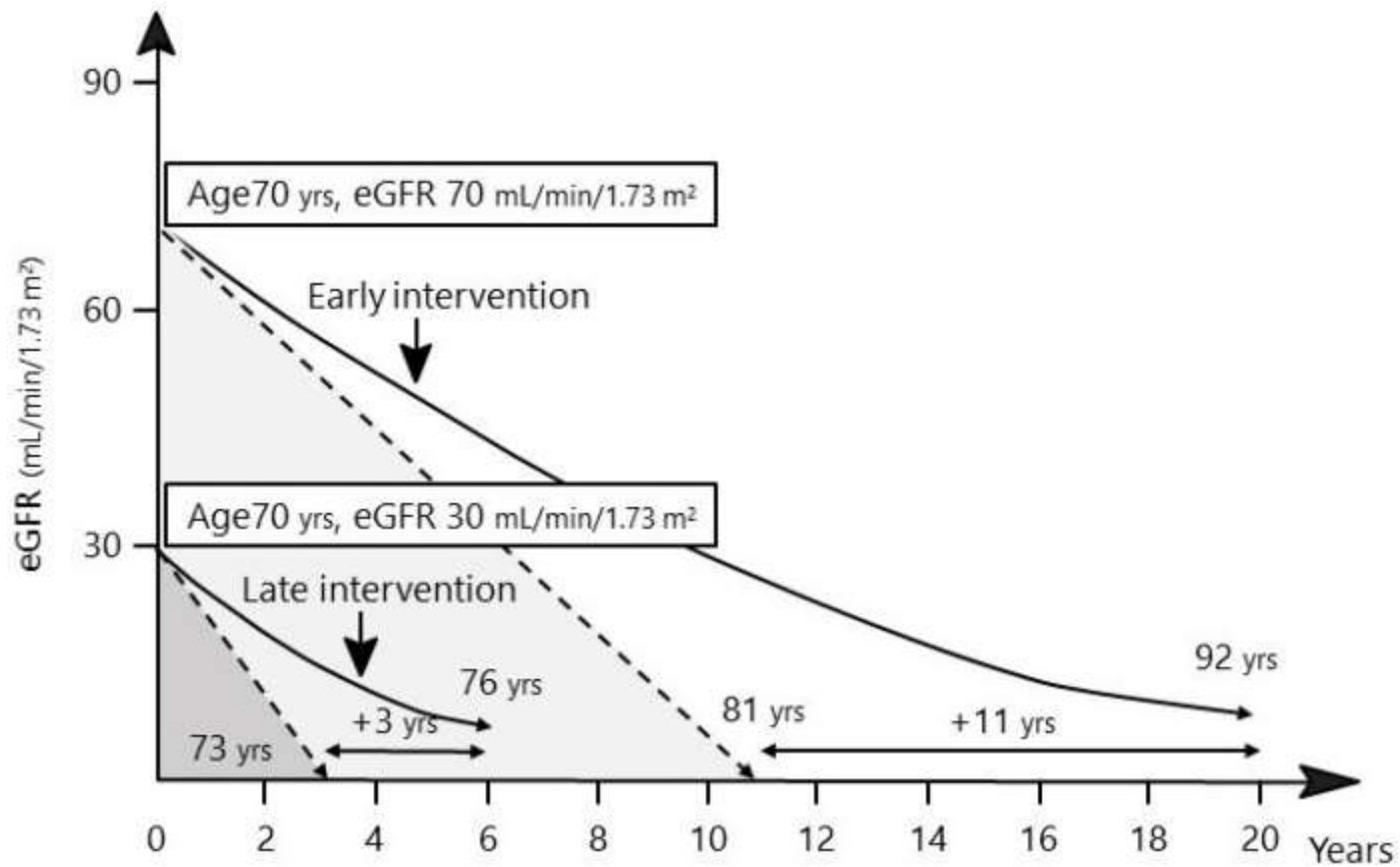





**No. of Participants**

Placebo	2152	2029	1981	1866	1795	1753	1672	1443	935	447	157
Dapagliflozin	2152	2031	2001	1896	1832	1785	1705	1482	978	496	157


Dapagliflozin in Patients with CKD NEJM Oct 2020



# NICE CKD Guidelines

- Screening at risk groups
  - Kidney Failure Risk Equation instead of eGFR threshold for referral
  - Statin treatment
  - SGLT2i
- 

# Risk factors for CKD

- Nephrotoxic medication (annually)
  - Diabetes
  - Hypertension
  - AKI
  - CVD
  - Structural renal tract disease, recurrent renal calculi or prostatic hypertrophy
  - Multisystem diseases with renal involvement
  - Gout
  - FHx ESKD or hereditary kidney disease
  - Incidental haematuria or proteinuria
- 

# THE KIDNEY FAILURE RISK EQUATION

Find out your real risk of kidney failure

[KIDNEY FAILURE RISK CALCULATOR](#) [LEARN HOW TO TEST YOUR KIDNEYS](#)

## FACTS & FIGURES OF CHRONIC KIDNEY DISEASE (CKD)

### CKD STAGES

Your kidneys primary function is to filter and excrete waste products. To find out how well your kidneys are doing, we measure the quantity of waste that circulates in your blood.

### eGFR ESTIMATED GLOMERULAR FILTRATION RATE

This is a test used to check how well the kidneys are working by estimating how much waste is in your blood. The more waste products in your blood the lower the filtration rate.

STAGE	eGFR (%)
1	> 90%
2	90-100%
3	60-89%
4	30-59%
5	< 15%

## KIDNEY FAILURE RISK CALCULATION

If you don't have the information required below talk to your doctor.

Age (Yrs)

Sex

eGFR (ML/MIN/1.73M<sup>2</sup>)

Urine Albumin: Creatinine Ratio

[SUBMIT](#)

<https://kidneyfailurerisk.co.uk/>



# YOUR RESULTS

 **88** mg/mmol  
URINE ALBUMIN

 **M**  
SEX

 **42**  
AGE

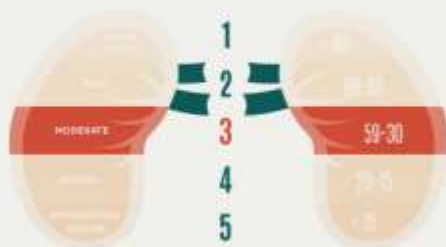
 **45** mL/min/1.73 m<sup>2</sup>  
EGFR

## ASSESSMENT

### STAGE 3

MODERATE DECREASE IN FUNCTION

CKD STAGES



ESTIMATED GLOMERULAR  
FILTRATION RATE

Patient risk of progression to kidney failure requiring dialysis or  
transplant:

AT 2 YEARS

1.9 %

AT 5 YEARS

6.5 %

#### NICE Referral Criteria:

Taking into account the individual's wishes and other health conditions, considering referral to a hospital kidney doctor if:

- 5-year KFRE predicted risk over 5% risk KFRE

## HOW CAN I REDUCE MY RISK OF KIDNEY FAILURE?

There are things you can do to reduce your risk of kidney failure over the next five years. Click below to see how the following will decrease your risk.



- Your current 5 year risk based on the answers you provided is **6.51%**
- Achieving good blood pressure control can reduce your 5 year risk from **6.51%** to **5.14%**.
- An ACE inhibitor (pril) or ARB (sartan) can reduce your 5 year risk from **6.51%** to **4.56%**.
- An SGLT2 inhibitor (gliflozin) can reduce your 5 year risk from **6.51%** to **3.58%**.

**The benefits of these changes can add up over time.**

## NENC Guideline For Use Of Sodium Glucose Co-transporter 2 Inhibitors (SGLT2i) in Adults

### Introduction & Scope

Aims of this document:

- To guide prescribing of SGLT2 inhibitors within each individual drug's current license
- To advise on appropriate choice of SGLT2 inhibitor
- To ensure safe and appropriate prescribing of SGLT2 inhibitors for patients
- To ensure that the necessary safety information is given to all patients

*It is important to note this is only a guide and not exhaustive, appropriate clinical judgement and referral to other reference sources may be appropriate in individual patient cases. This resource is only for use in adults.*

*Licenses for SGLT2 inhibitors are changing rapidly. Always check the up-to-date licenses.*

*The information in this guidance was correct at the time of publication.*

### What are SGLT2 inhibitors?

- An established class of medications which are licensed for the treatment of:
  - Insufficiently controlled type 2 diabetes (T2DM)
  - Symptomatic chronic heart failure (HF) with reduced ejection fraction (HFrEF), with mildly reduced ejection (HFmrEF), or with preserved ejection fraction (HFpEF) (with or without T2DM)
  - Chronic kidney disease (CKD) (with or without T2DM)
- SGLT2 inhibitors have been shown to reduce the risk of cardiovascular events in people living with T2DM and atherosclerotic cardiovascular disease (ASCVD) i.e., coronary heart disease, acute coronary syndrome, previous myocardial infarction, stable angina, prior coronary or other revascularisation, cerebrovascular disease (ischaemic stroke and transient ischaemic attack) and peripheral arterial disease
- They act by preventing the absorption of glucose and sodium, mainly from the proximal renal tubule in the kidney
- Glucose and sodium are, therefore, lost in urine
- People do not become hyponatraemic (unless co-prescribed diuretics) as most of the sodium is reabsorbed in the distal tubule
- This effect results in decreasing the blood glucose level, weight loss, an osmotic diuresis and a drop in blood pressure

### Step 1: Identify if the person is suitable for an SGLT2 inhibitor – decide which SGLT2 inhibitor to use

#### Which SGLT2 inhibitor should I use?

Choice of SGLT2 inhibitor depends on co-morbidities.

##### One significant co-morbidity:

T2DM	HFrEF	HFpEF or HFmrEF	CKD*
Dapagliflozin or Empagliflozin	Dapagliflozin or Empagliflozin	Dapagliflozin	Dapagliflozin

##### Two significant co-morbidities:

	T2DM	HFrEF	HFpEF or HFmrEF	CKD*	CKD**	ASCVD***
T2DM		Dapagliflozin or empagliflozin	Dapagliflozin	Dapagliflozin	Dapagliflozin	Empagliflozin or dapagliflozin
HFrEF	Dapagliflozin or empagliflozin			Dapagliflozin	Dapagliflozin	Empagliflozin or dapagliflozin
HFpEF or HFmrEF	Dapagliflozin			Dapagliflozin	Dapagliflozin	Dapagliflozin
CKD*	Dapagliflozin	Dapagliflozin	Dapagliflozin			Dapagliflozin
CKD**	Dapagliflozin	Dapagliflozin	Dapagliflozin			
ASCVD***	Empagliflozin or dapagliflozin	Empagliflozin or dapagliflozin	Dapagliflozin	Dapagliflozin		

##### Three or more significant co-morbidities (T2DM and/or ASCVD and/or HF and/or CKD\*):

Dapagliflozin (or empagliflozin if no CKD)

We would not advocate switching between SGLT2 inhibitors if co-morbidity changes.

The above suggestions for initial therapy are based on licences and clinical trial data including cardiovascular outcome trials.

##### Criteria For Use:

## SGLT2-inhibitors and Chronic Kidney Disease - Patient Information Leaflet



### What about driving?

Inform the DVLA if your doctor has told you that you are at risk of low blood sugar or if you experience low blood sugar.



### Do I need to be monitored?

Specific monitoring after starting an SGLT2 inhibitor is not required.

Your medical team will continue to monitor your kidney function as part of your routine care.

Unless you feel unwell, there is usually no need to have extra blood tests after starting an SGLT2 inhibitor.

Northern Treatment Advisory Group  
<https://ntag.nhs.uk>



### What are SGLT2 inhibitors?

SGLT2 inhibitors stands for Sodium Glucose Co-transporter-2 inhibitors. These drugs are sometimes also called 'gliflozins' or 'flozins' based on their naming. For example, DAPAGLIFLOZIN, CANAGLIFLOZIN and EMPAGLIFLOZIN are all SGLT2 inhibitors. SGLT2 inhibitors act on the kidney, causing loss of salt and sugar from the body.

### How do I take it?

The medicine is taken once daily, at the same time each day, with or without food. If you miss a tablet and remember later in the day you can take it. If you remember the following day, do not take two tablets together.



### Why should I be prescribed an SGLT2 inhibitor?

These drugs are used to treat diabetes, heart failure, and chronic kidney disease (CKD). Many people will feel no day-to-day difference from taking SGLT2 inhibitors. Large medical studies have shown for many people with CKD, type 2 diabetes or heart failure taking SGLT2 inhibitors alongside other medications can improve life expectancy and reduce risk of their condition getting worse.

Studies of SGLT2 inhibitors were undertaken in people already taking certain blood pressure medications. It is therefore likely that your Doctor will check that you are taking the correct doses of these medications, even if your blood pressure is normal, before starting an SGLT2 inhibitor.

# Q&A session

- Any questions?

# Upcoming events...

# Using digital systems to detect and manage patients for CKD

23rd April 2024 12.00-13:00



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information

