

# Starting Dialysis means that medical care has failed:

- 73-year-old, male with type II diabetes
- End stage kidney disease (diabetic nephropathy)
- Haemodialysis 3x weekly
- Retinopathy (registered blind)
- Previous non-STEMI
- Ejection fraction 40%
- Left mid foot amputation
- Wheelchair



# Making a diagnosis of CKD



## Blood test

- Creatinine based GFR estimation (eGFR)
- Calculation needs creatinine, age, gender
- Result expressed as  $\text{ml}/\text{min}/1.73\text{m}^2$
- No ethnicity correction



## Urine test

- Any urine
- Request albumin:creatinine ratio (UACR)
- Expressed as  $\text{mg albumin}/\text{mmol creatinine}$
- (Dipstix less accurate)

# CKD classification by eGFR and albuminuria

eGFR ml/min/1.73m <sup>2</sup>	Albuminuria categories (Based on urinary albumin:creatinine ratio)		
	A 1 <3 mg/mmol	A 2 3-30 mg/mmol	A 3 >30 mg/mmol
G1 ≥ 90	No CKD	G1 A2	G1 A3
G2 60-89	No CKD	G2 A2	G2 A3
G3a 45-59	G3a A1	G3a A2	G3a A3
G3b 30-44	G3b A1	G3b A2	G3b A3
G4 15-29	G4 A1	G4 A2	G4 A3
G5 <15	G5 A1	G5 A2	G5 A3

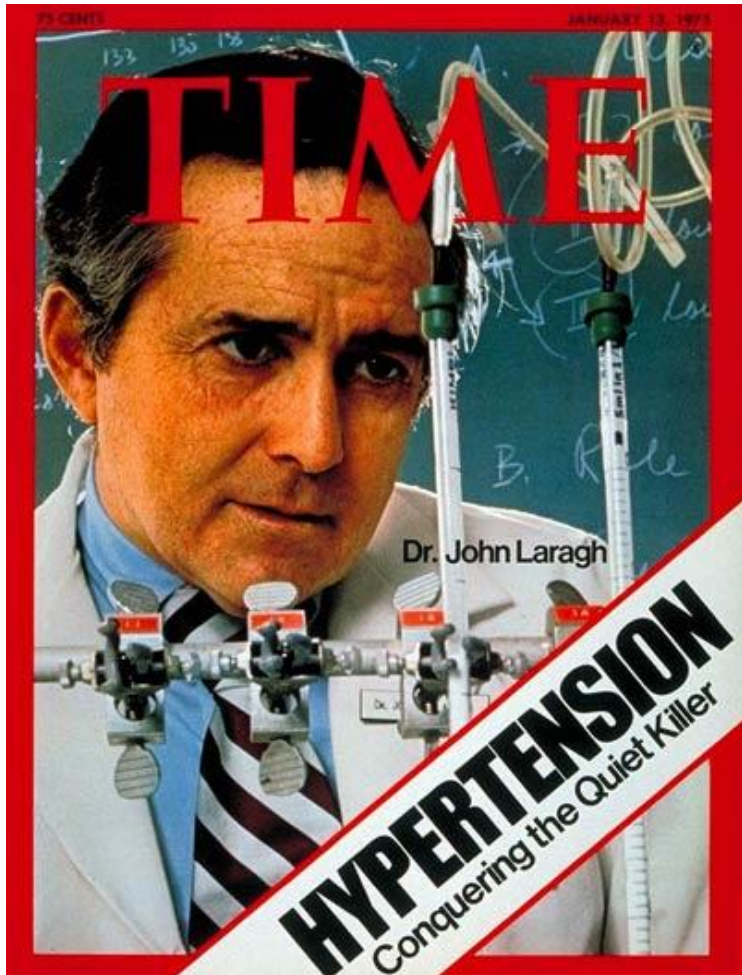


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# Should persons with hypertension, diabetes, and cardiovascular disease should be screened for CKD?



# Seek and you will find

## ANALYSIS

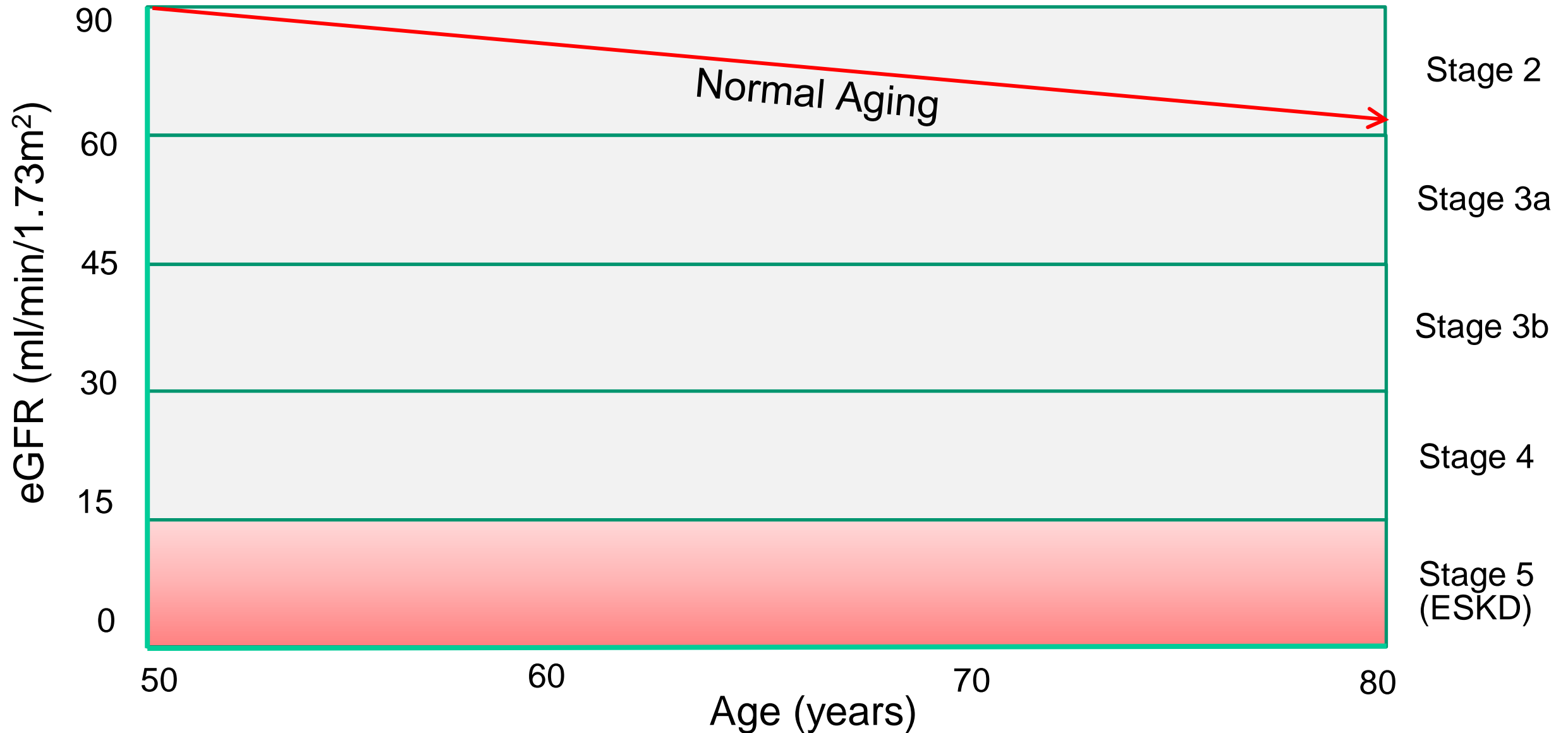
# Re-evaluating national screening for chronic kidney disease in the UK

New drugs have increased the potential benefits of screening but many unanswered questions remain about the best way to identify people for treatment, say **Vageesh Jain and colleagues**

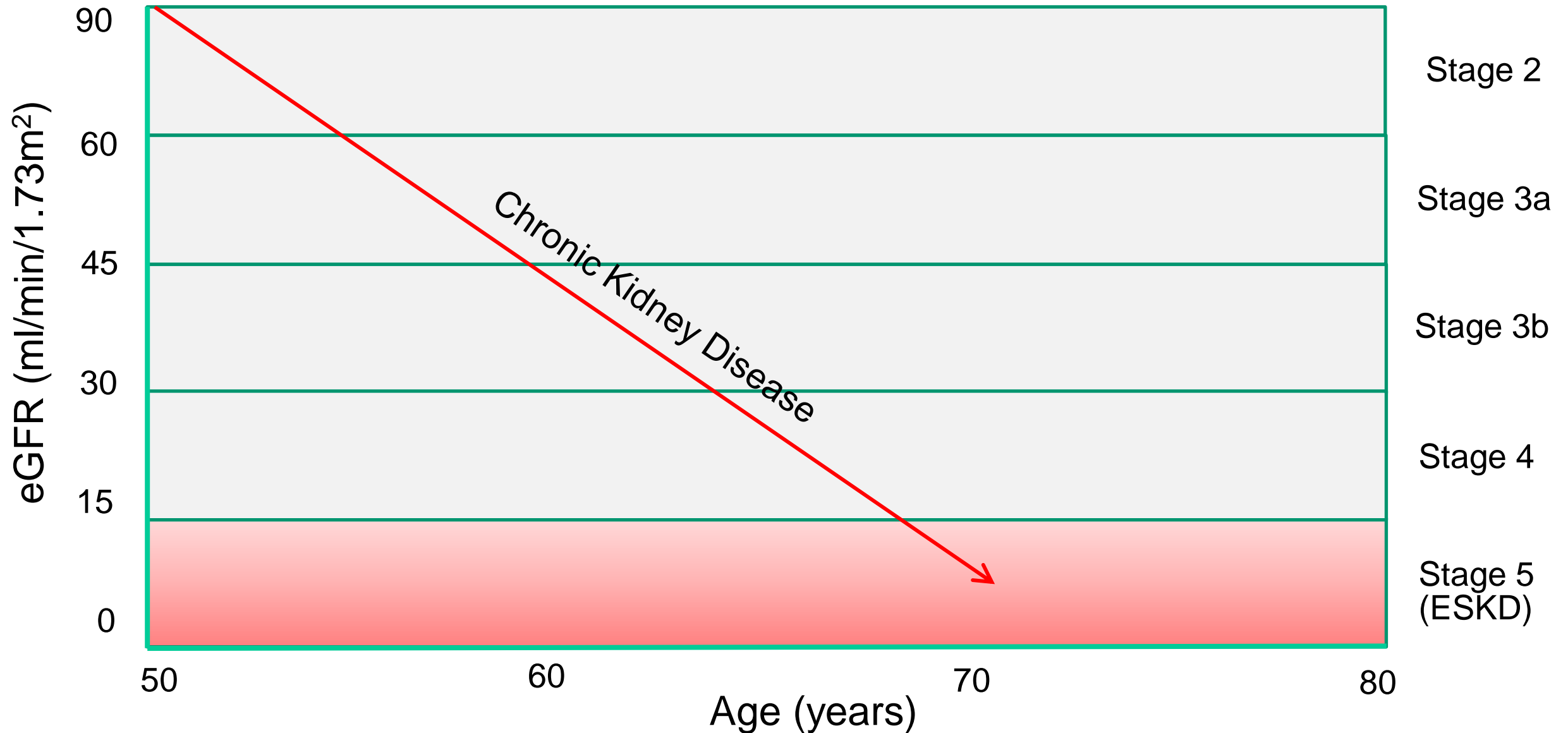


SATURN STILLIS/SPL

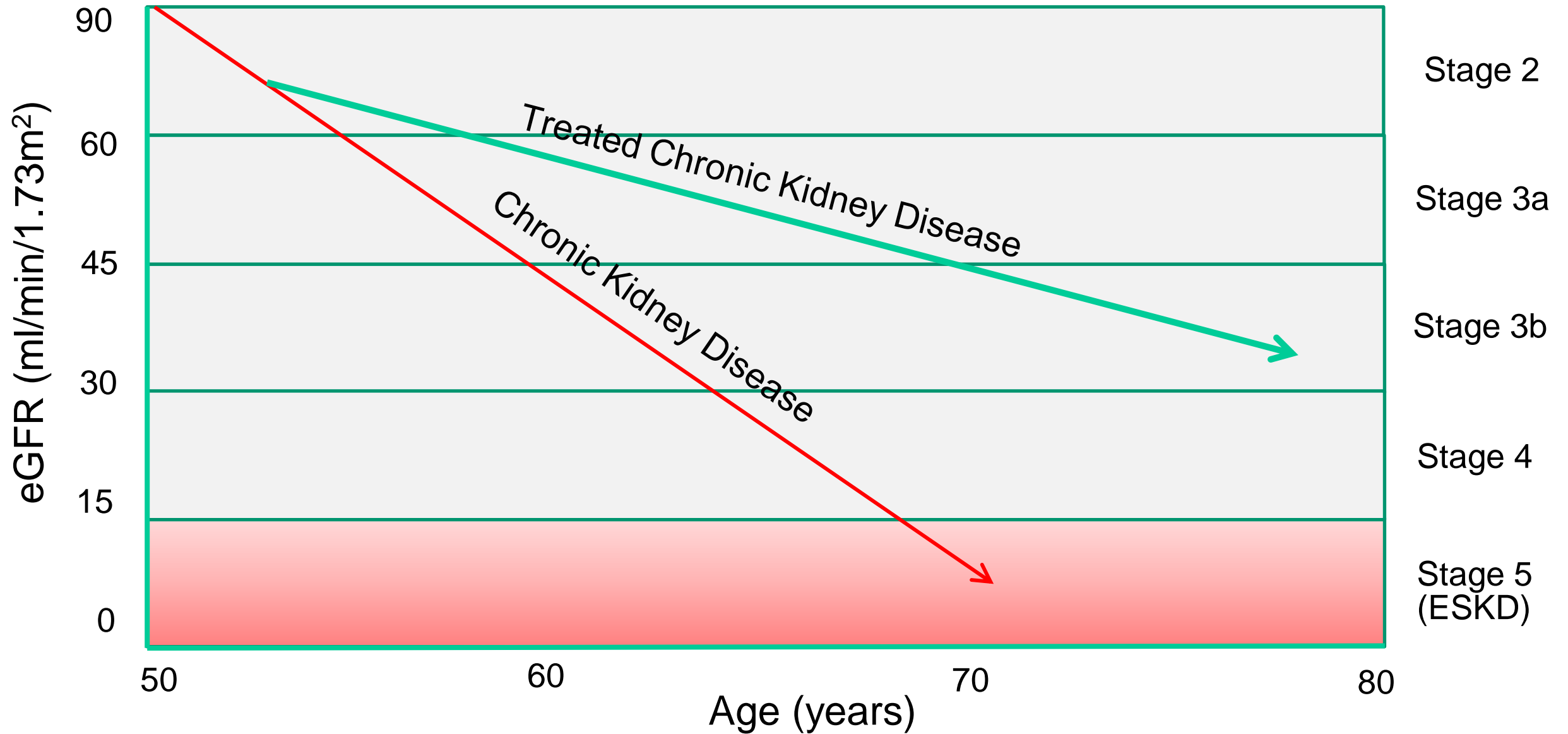
# Progressive decline in kidney function



# Progressive decline in kidney function



# Kidney Protection



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Increasing risk

Increasing risk

# Care 1: White male with Type 2 diabetes and CKD

- 68-year-old white man with type II diabetes for 10 years
- Weight 135 kg, BMI 31.8 kg/m<sup>2</sup>
- Early diabetic retinopathy
- Metformin 500 mg bd, Gliclazide 40 mg bd
- Ramipril 10 mg od, Atorvastatin 20 mg od
- Annual review
- BP 155/95 mmHg

eGFR 52 ml/min/1.73m<sup>2</sup> (previously 58)

UACR 672 mg/mmol (previously 309)

HbA1c 72 mmol/mol (8.7%)



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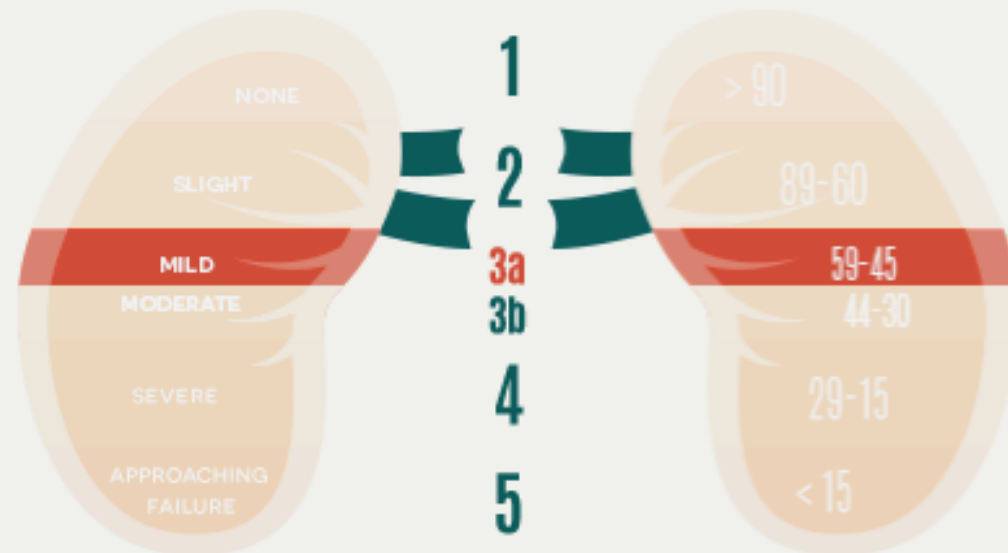
# STAGE 3a

MILD DECREASE IN FUNCTION

CKD STAGES

ESTIMATED GLOMERULAR  
FILTRATION RATE

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



AT 2 YEARS

AT 5 YEARS

1.2%

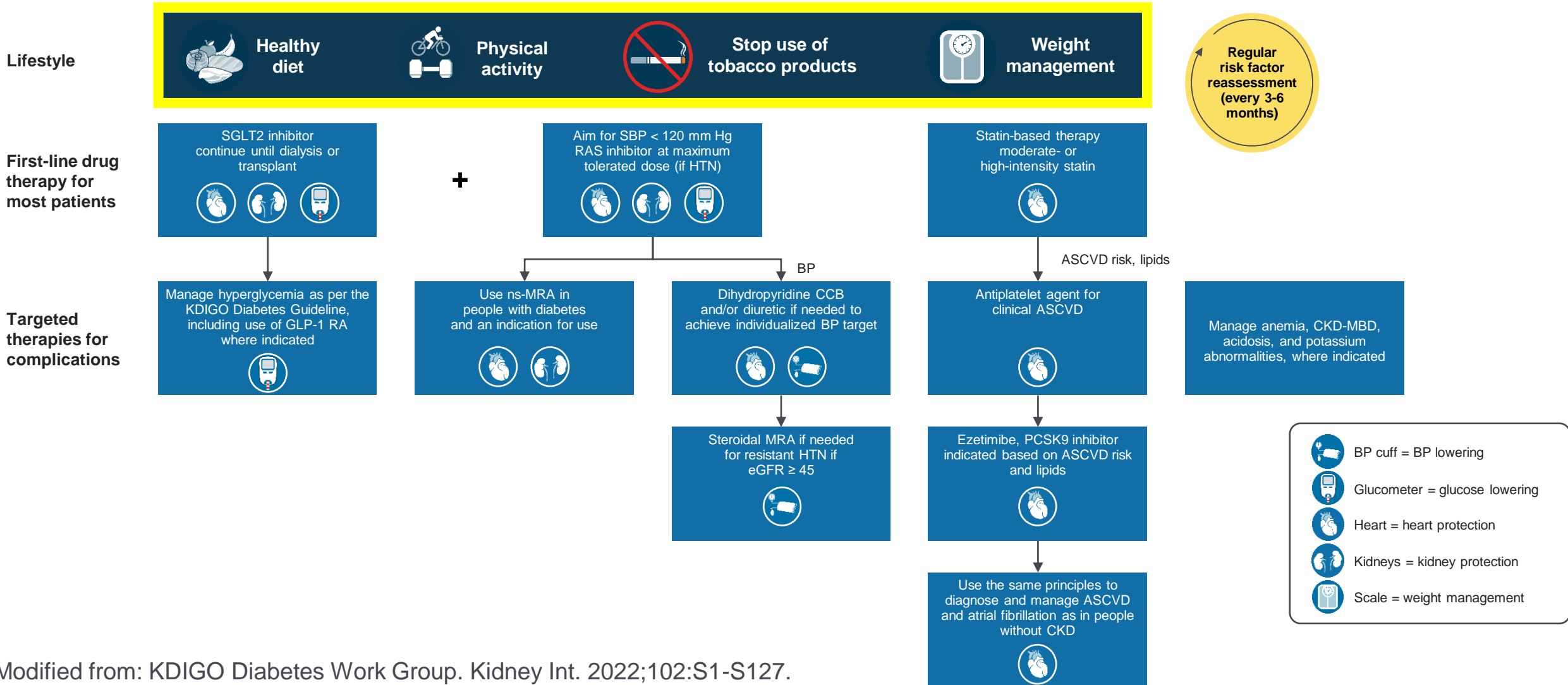
4.3%

## NICE Referral Criteria:

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

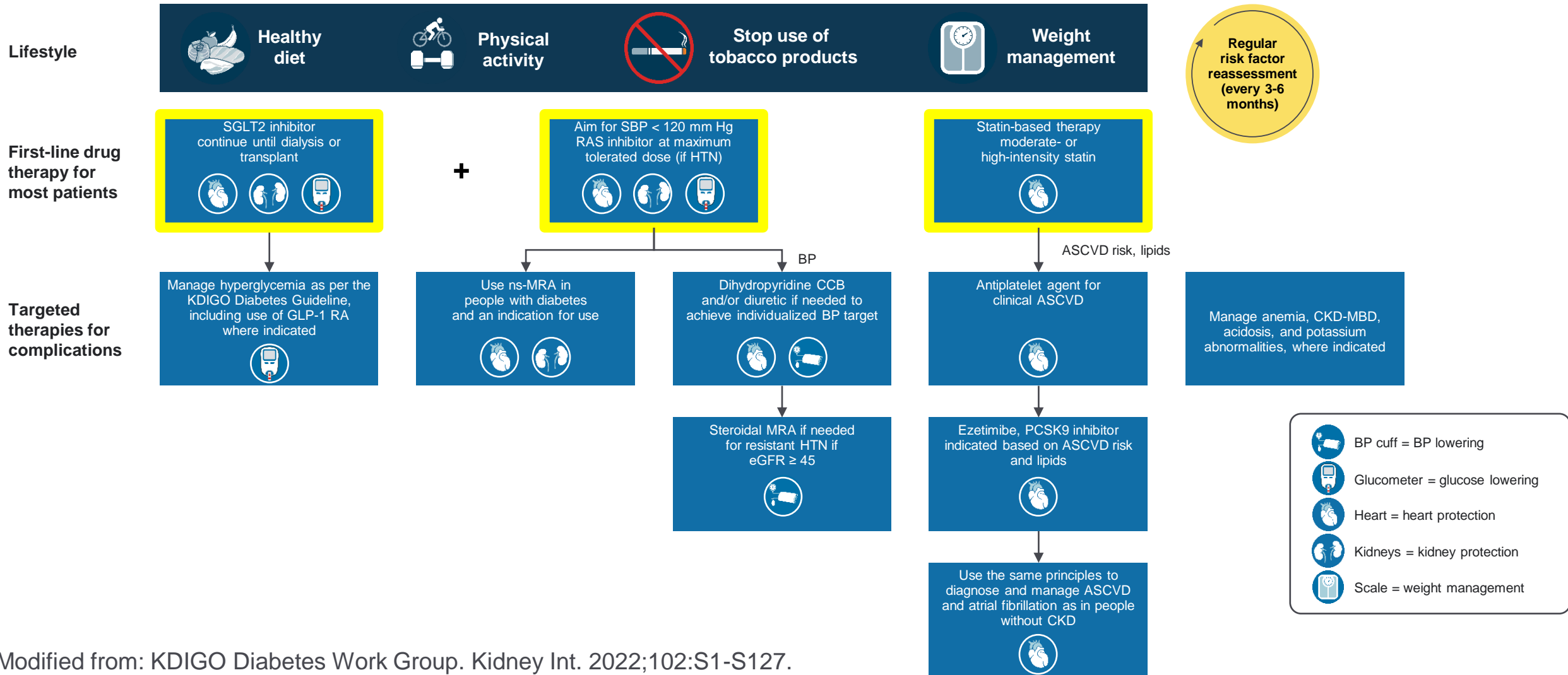
- 5-year KFRE predicted risk over 5% risk KFRE

# 2024 KDIGO CKD Evaluation and Management Guideline



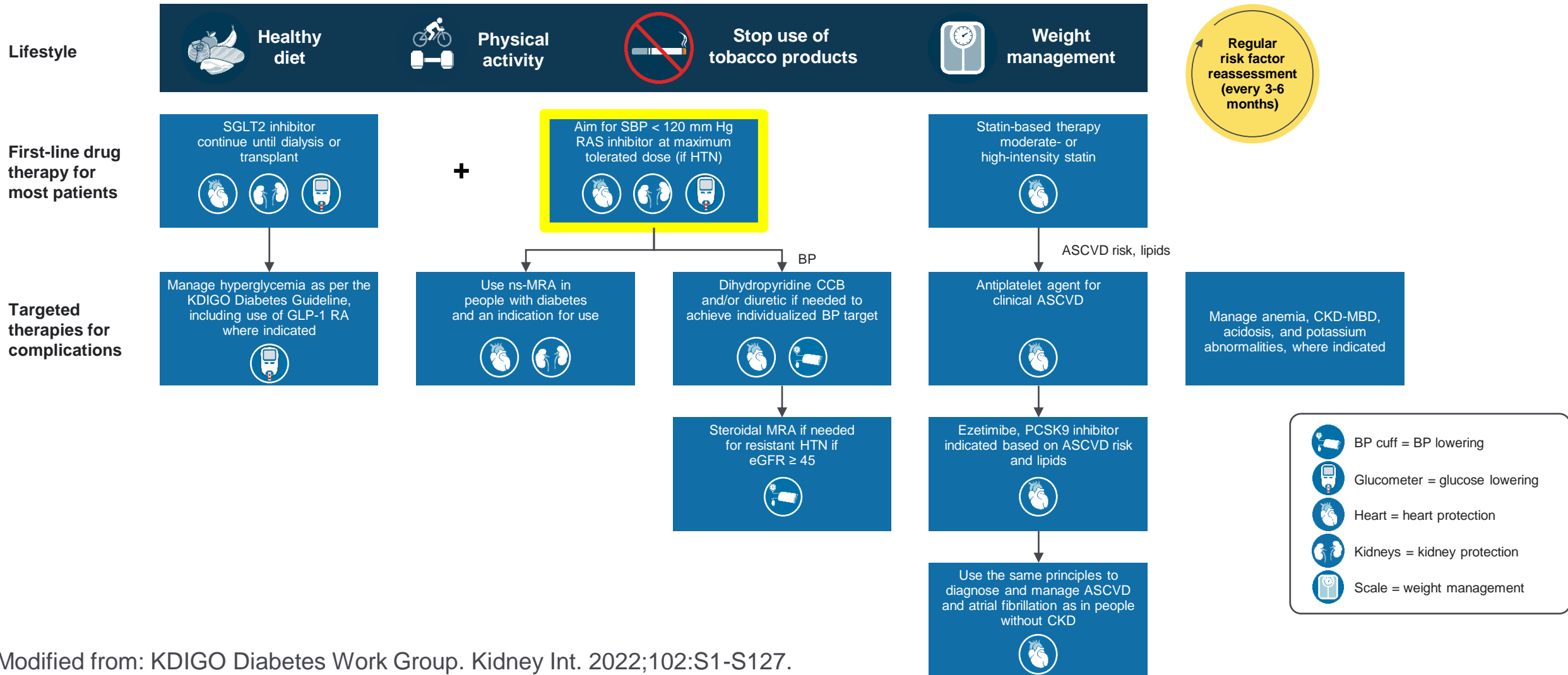
Modified from: KDIGO Diabetes Work Group. *Kidney Int.* 2022;102:S1-S127.  
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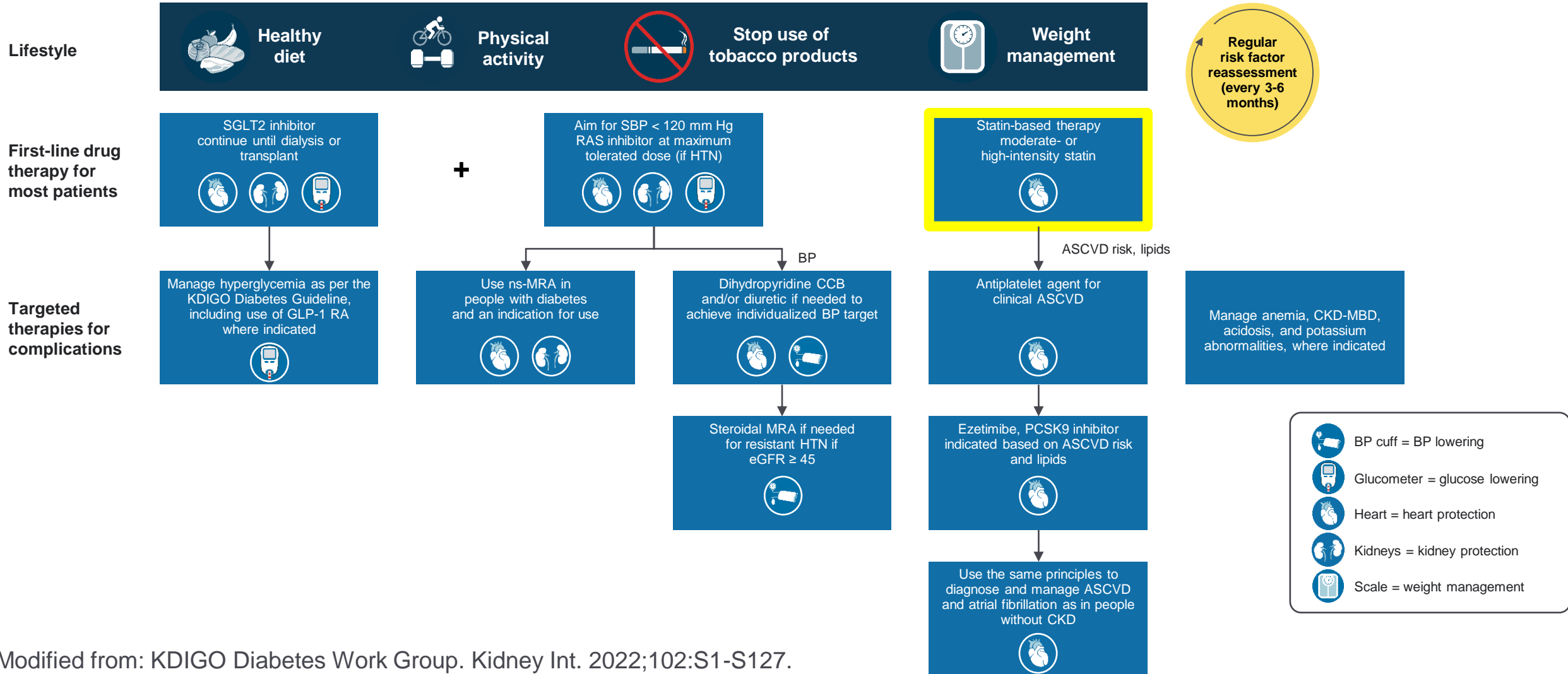
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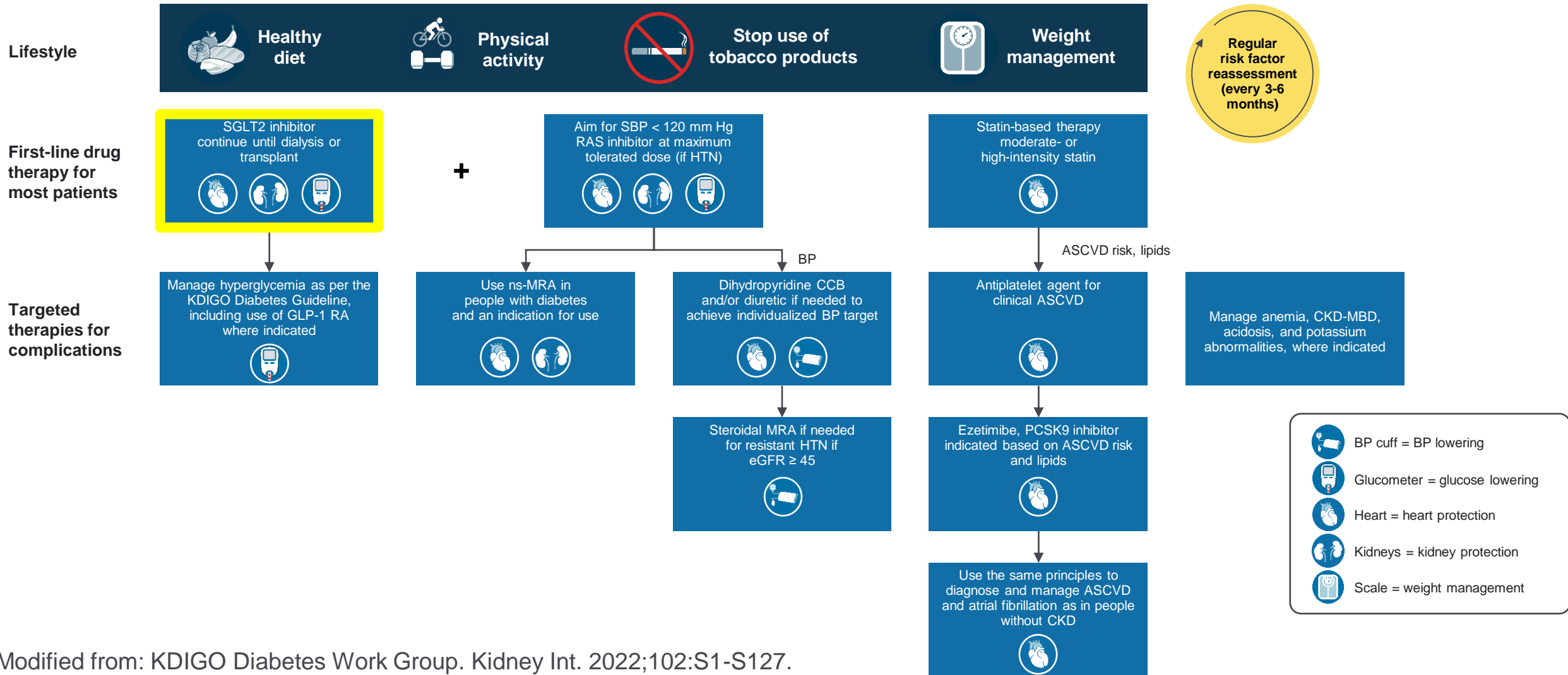
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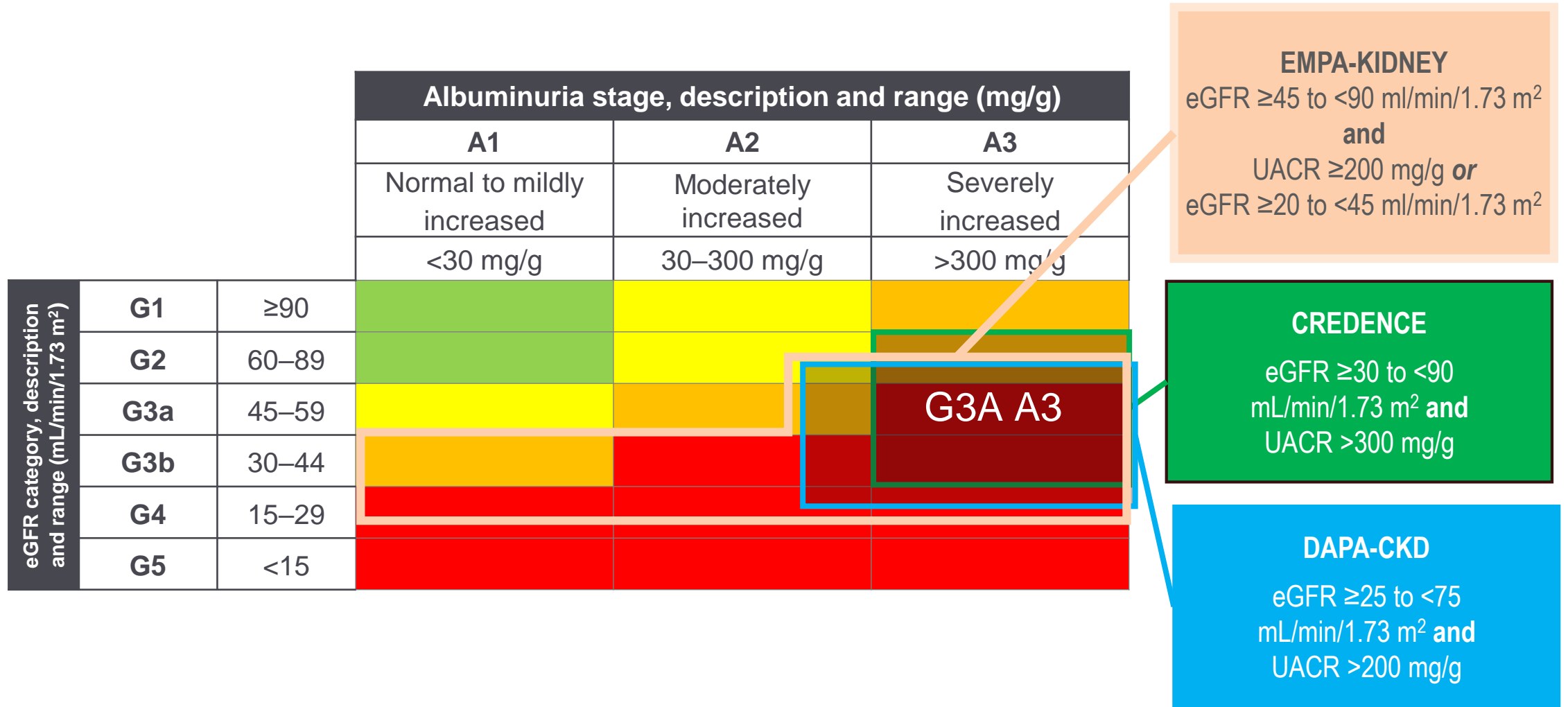
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# Comparison of CREDESCENCE, DAPA-CKD and EMPA Kidney Trials



CKD, chronic kidney disease; eGFR, estimated glomerular filtration rate; UACR, urine albumin-to-creatinine ratio

The EMPA-KIDNEY Collaborative Group. *Nephrol Dial Transplant* 2022;gfac040; Kidney Disease: Improving Global Outcomes (KDIGO) CKD Work Group. *Kidney Int Suppl* 2013;3:1; Perkovic V *et al. N Engl J Med* 2019; 380:2295; Wheeler DC *et al. Nephrol Dial Transplant* 2020;35:1700

# Inclusion Criteria: SGLT2 inhibitor trials in Chronic Kidney Disease

	<b>CREDENCE<sup>1</sup></b>	<b>DAPA-CKD<sup>2</sup></b>	<b>EMPA-KIDNEY<sup>3</sup></b>
<b>Published</b>	April 2019	September 2020	November 2022
<b>SGLT2 inhibitor</b>	Canagliflozin	Dapagliflozin	Empagliflozin
<b>Non diabetic patients</b>	No	Yes	Yes
<b>eGFR range</b>	30 to 90 ml/min/1.73m <sup>2</sup>	25 to 75 ml/min/1.73m <sup>2</sup>	20 to 90 ml/min/1.73m <sup>2</sup>
<b>UACR range</b>	300-5000 mg/g 33.9-565 mg/mmol	200-5000 mg/g 22.6-565 mg/mmol	> 200 mg/g (22.6 mg/mmol) if eGFR >45 ml/min/1.73m <sup>2</sup>

<sup>1</sup>Perkovic V et al, NEJM 2019;380:2295-2306

<sup>2</sup>Heerspink HJL et al, NEJM 2020;383:1436-1446

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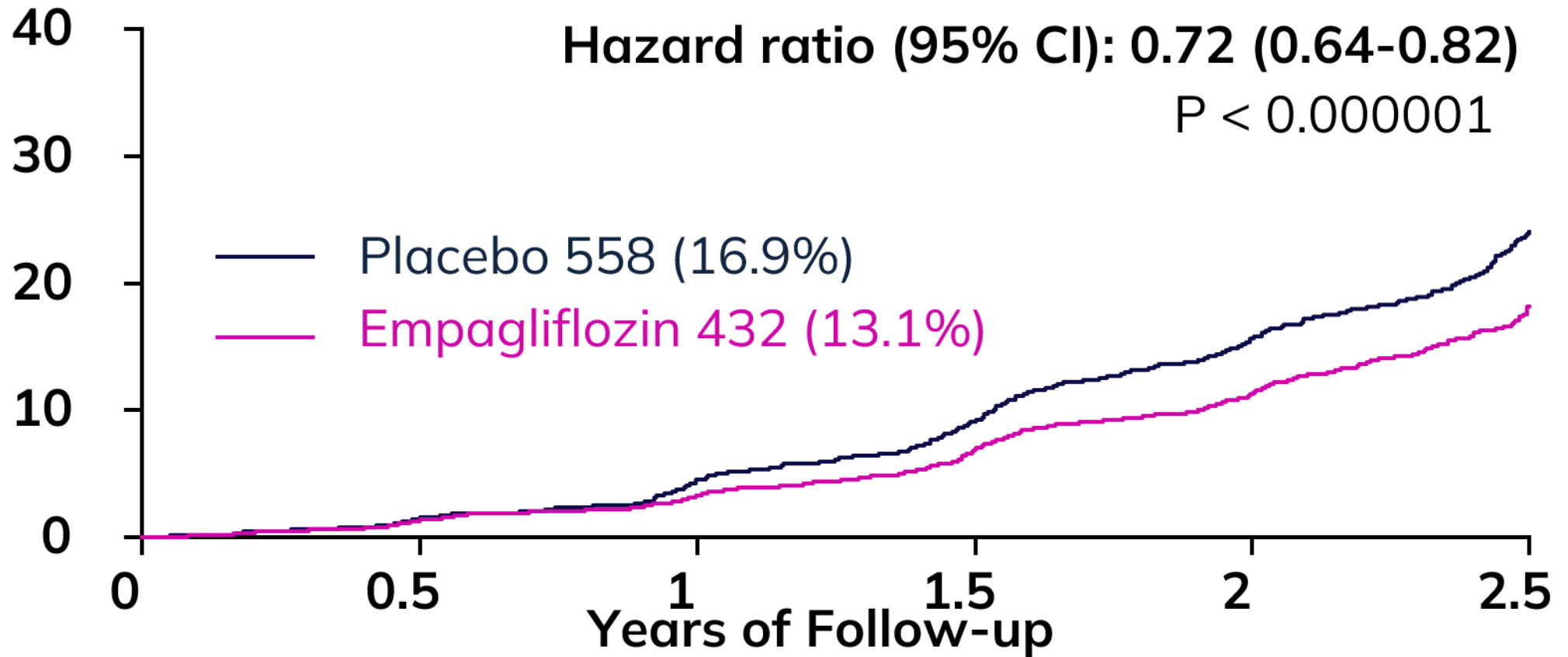
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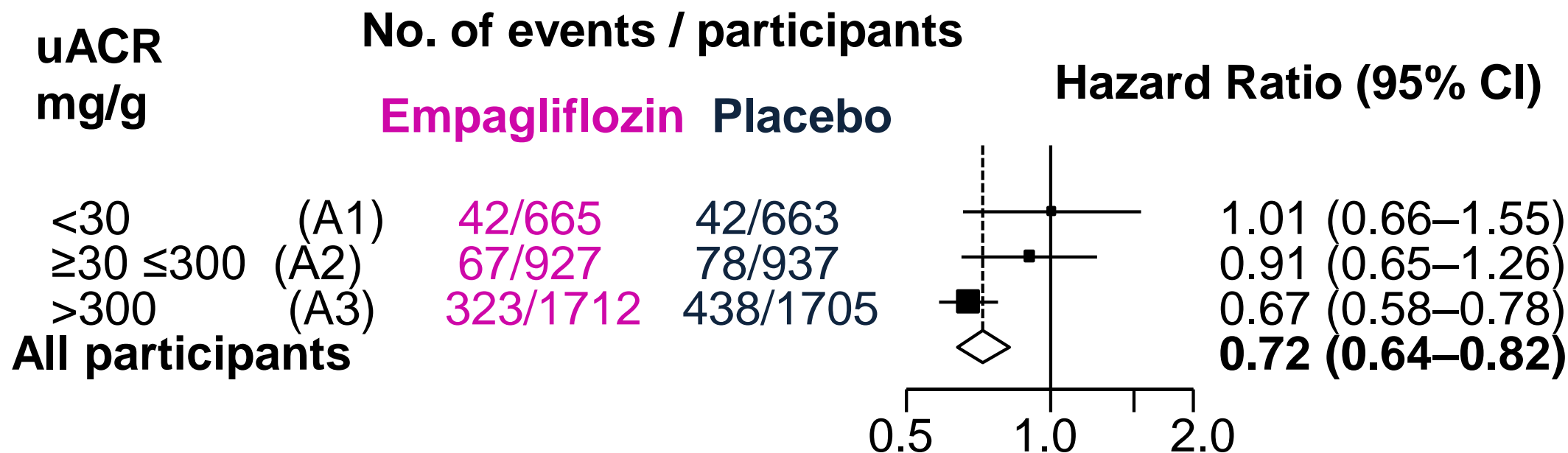
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# EMPA Kidney: Primary Composite Outcome

(CV death, or ESKD, >40% eGFR decline or to <10 ml/min/1.73m<sup>2</sup> or renal death)



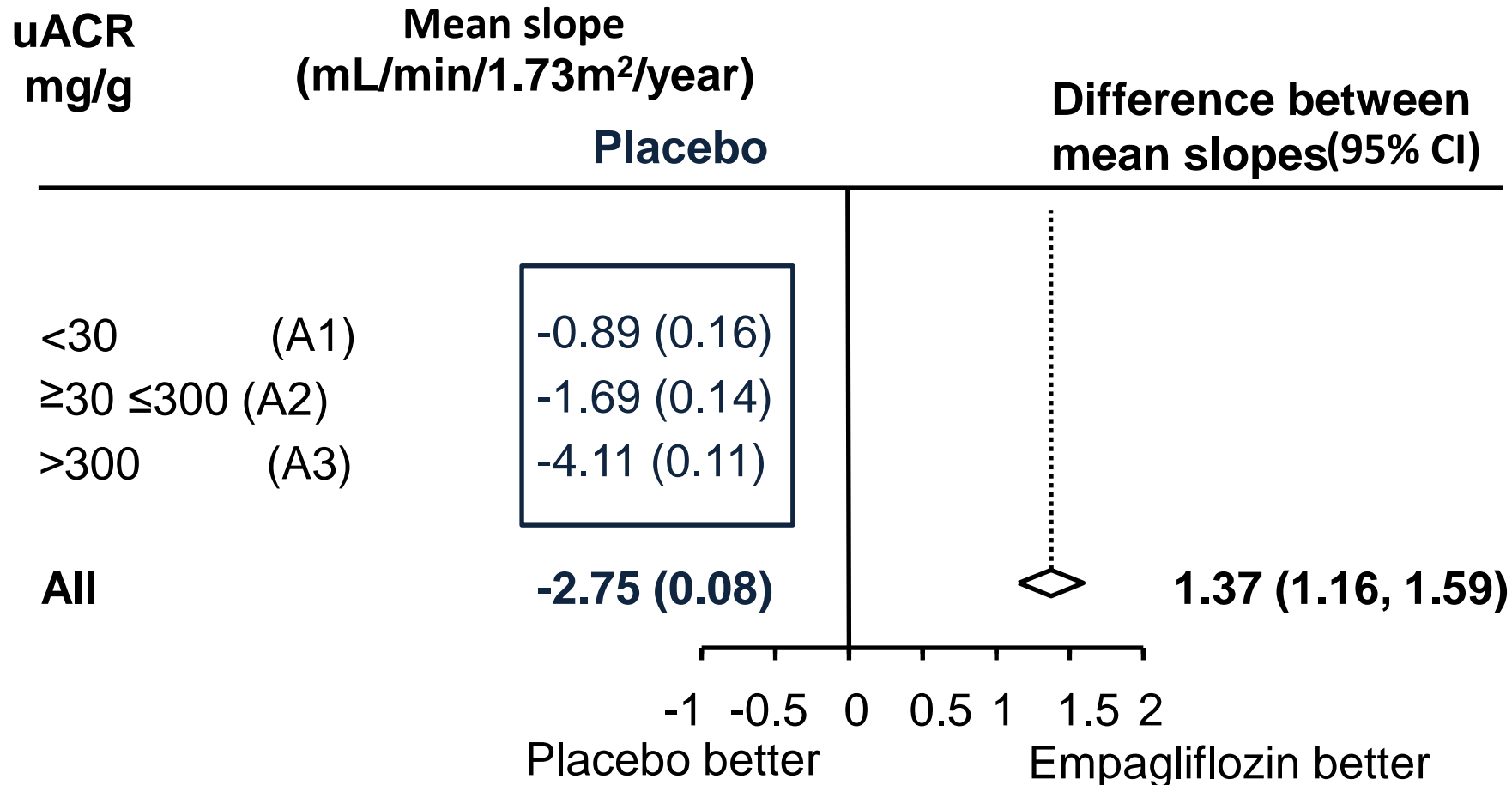
# EMPA Kidney: Primary outcome by albuminuria



Trend P value= 0.02

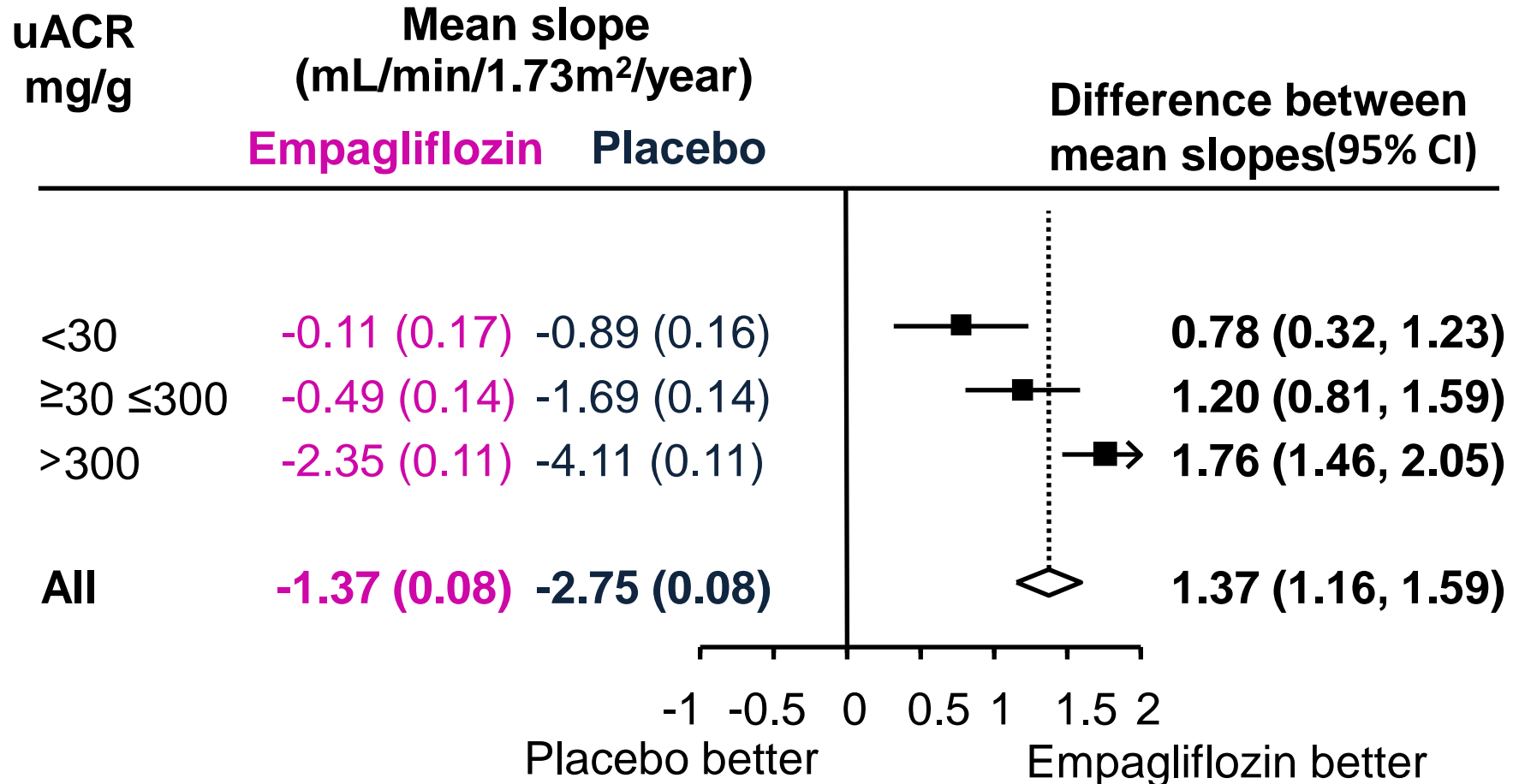
# EMPA Kidney: Chronic eGFR slopes by albuminuria

Prespecified exploratory analyses



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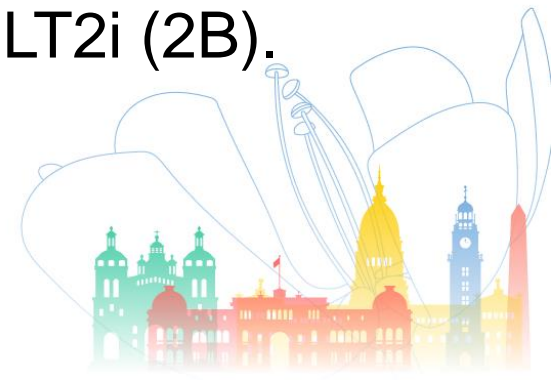
# TAKE HOME MESSAGES: SGLT2 INHIBITORS IN CKD

In which patients should SGLT2 inhibitors be considered?

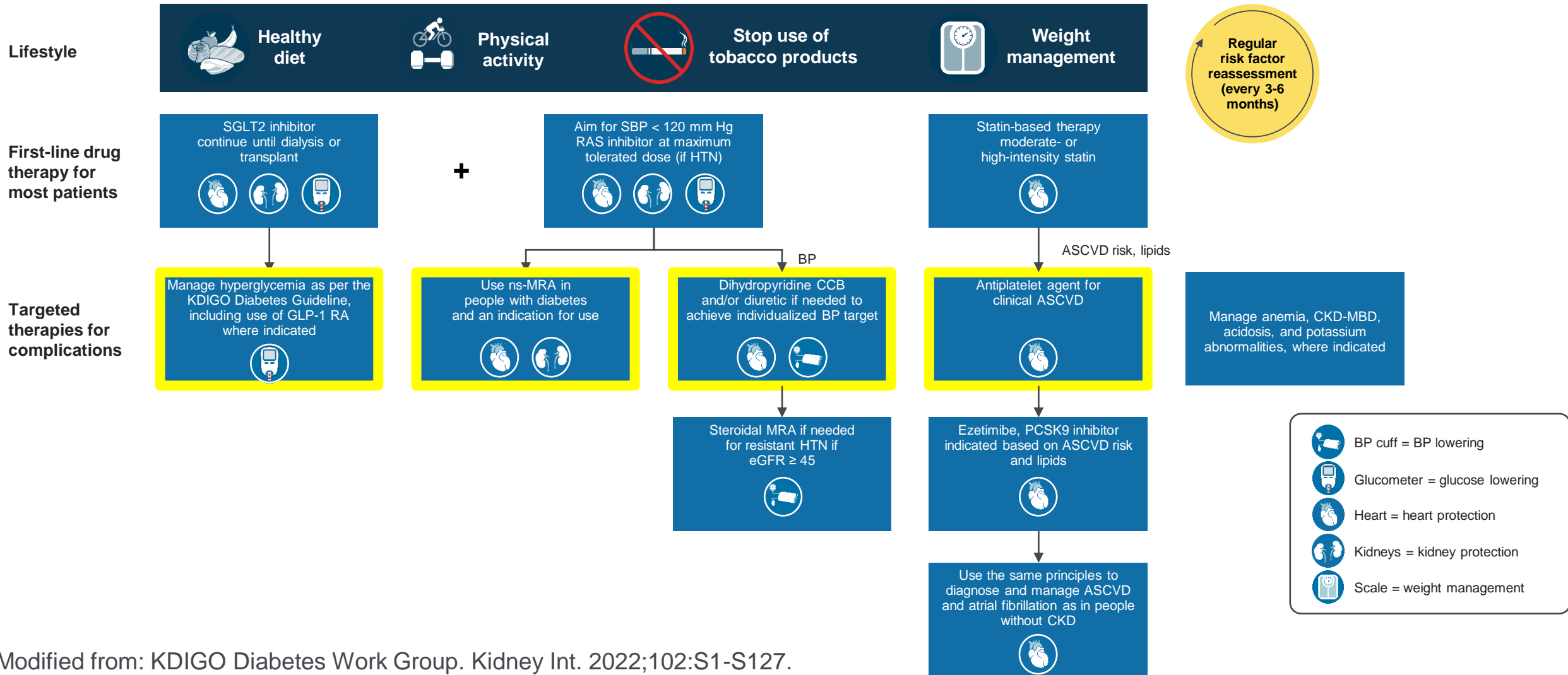
**Recommendation 3.7.1:** We recommend treating patients with type 2 diabetes (T2D), CKD, and an eGFR  $>20$  ml/min per  $1.73$  m<sup>2</sup> with an SGLT2i (1A).

**Recommendation 3.7.2:** We recommend treating adults with CKD with an SGLT2i for the following (1A): eGFR  $>20$  ml/min per  $1.73$  m<sup>2</sup> with urine ACR  $>200$  mg/g ( $>20$  mg/mmol), or heart failure, irrespective of level of albuminuria.

**Recommendation 3.7.3:** We suggest treating adults with eGFR 20 to 45 ml/min per  $1.73$  m<sup>2</sup> with urine ACR  $<200$  mg/g ( $<20$  mg/mmol) with an SGLT2i (2B).



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- Early diabetic retinopathy
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- Ramipril 10 mg od, Atorvastatin 20 mg od, Furosemide 40 mg od
- Dapagliflozin 10 mg od,
- Finerenone 25 mg od
- Semaglutide 2.4 mg weekly



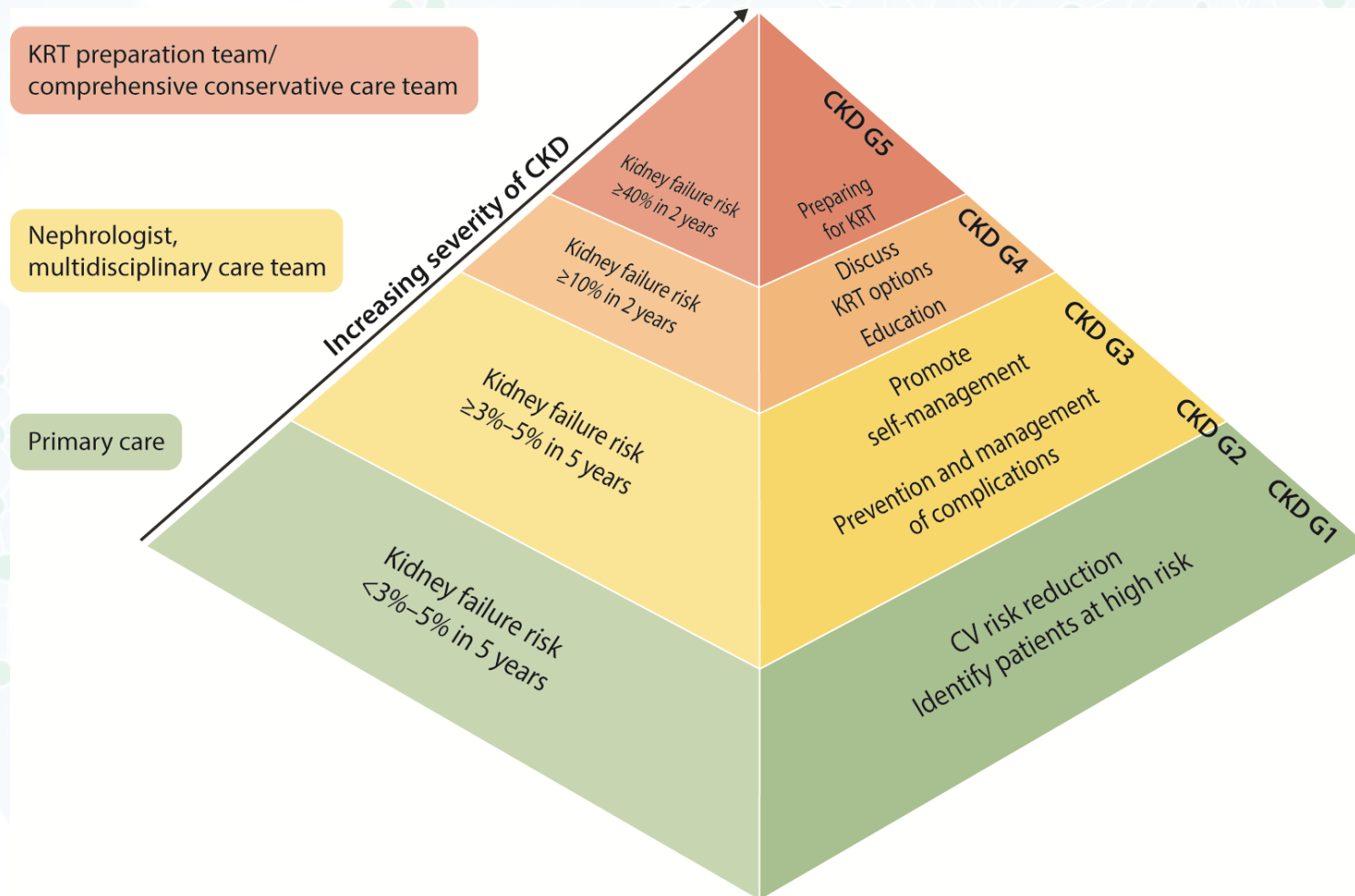
# Four Pillars of CardioRenal protection in Diabetic Kidney Disease



Concept borrowed from: Four pillars of heart failure: Contemporary pharmacological therapy for heart failure with reduced ejection fraction. Straw S. <http://orcid.org/0000-0002-2942-4574>

# MANAGEMENT – ADVANCED CARE PLANNING

Plans addressing future health care states should be jointly agreed with people with CKD and their families/carers and known to all. Advanced care planning for those choosing supportive care is particularly important.



# The talk in one slide

- Many recent developments in preventative nephrology which should improve patient outcomes and help avoid CV events dialysis in CKD.
- Lifestyle modifications, lowering blood pressure and controlling diabetes underpin management of CKD (?Primary Care).
- Statin-based regimens reduce the risk of atherosclerotic cardiovascular events in CKD patients (less benefit in ESKD).
- ACEi/ARB slow progression in patients with albuminuria
- SGLT2 inhibitors are renoprotective in CKD patients and reduce the risks of cardiovascular complications, particularly heart failure.
- Additional renoprotective therapies in patients with type 2 diabetes include 3<sup>rd</sup> Generation MRAs and GLP-1 RAs.

