

Diabetes Unfiltered: The Management of CKD & DKD

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Disclosures 2024/5

Speaker Fees: AstraZeneca, Bayer, Boehringer Ingelheim, Dexcom, Daiichi Sankyo, Grunenthal, Lilly, Menarini, Idorsia, Thornton & Ross, Boston Scientific

Consultancy Fees: AstraZeneca, Dexcom, Boehringer Ingelheim, Lilly, Menarini, Roche, Oviva, Idorsia, Grunenthal, Boston Scientific

Congress Attendance: Menarini, Daiichi Sankyo, Lilly, Bayer



ACR=albumin to creatinine ratio; AKI=acute kidney injury; BP=blood pressure; CKD=chronic kidney disease; CNI=calcineurin inhibitor; CVD=cardiovascular disease; CVE=cardiovascular event; eGR=estimated glomerular filtration rate; EPO=erythropoietin; ESRD=end=stage renal disease; GFR=glomerular filtration rate; KFRE=Kidney Failure Risk Equation; NSAID=nonsteroidal anti-inflammatory drug; NVH=non-visible haematuria; PKD=polycystic kidney disease; QoL=quality of life; RAG=red, amber; green; sCr=serum creatinine; uACR=urine albumin to creatinine ratio; UTI=urinary tract infection

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v this Primary Care Hack online at medscape-uk.co/Hack-CKD

Last updated: Februa



Disease in Primary Care

Medscape ***** UK X Guidelines

Primary Care Hacks

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	For primary prevention, start atorvastatin 20 mg OD	Lifest	yle and	Dietary Mo	dificatio	n	
1. Lipids ^[3–5,15–18]	 For secondary prevention, do not stop or attenuate dose if eGFR <30 m//min/1.73 m² Offer aspirin for secondary prevention of CVD See also the <u>Primary Care Hack on lipid management</u>. 	 Encour smokin Advise (ideally 	age weigh ng cessatio on salt re <2 g of so	nt loss and on ^[3-6,8] striction odium per	with incre and prog • Avoid NS • Promote	ased risk of n ression of CKI AIDs ⁽³⁻⁵⁾ exercise of at	ephriti D ^(13,14) t least
	• Aim for SBP <130 mmHg , or as low as reasonably achievable ^(A) • Follow <u>NG136⁽¹⁹⁾ and NG203^[0] when choosing medications</u>	day, ec sodium • If gastr consid	uating to chloride) ^[] ic protection er H_RAs c	<5 g of 15.8,121 on is required, over PPIs, as	 150 minu Kidney Ca a helpful leaflet to 	tes per week ¹² are UK has pro patient inform support peop	aduce
	When RAASi is first line, choose an ARB ^[8] Independent of BP: ⁽⁸⁾ if uACR >30 mo/mmol: start ARB and titrate to	PPI use	has been	associated	with CKD		
2. RAASis	maximum tolerated dose	riy	ure 1. ivia	Related to F	AASis ^{[3,22-3}		cuon
and Blood	start ARB and titrate to maximum tolerated dose			Seru	m K+ (mmo	I/I)	
Pressure	additional agents may be required			5.5–6.1	6.3	2-6.4	≥6.
	 If GURK <60 m/mm/1, 1.3 m² when starting or increasing KAASI, check U&E whith 28 days; If GCFR >60 m/min/1.73 m², there is no need to routinely recheck U&E If K+ 25.5 mmol/l or sCr rise 250% encountered, follow Figure 1; otherwise, continue to optimise RAASi dose. 	Unwell and/or ≥50% sCr rise	Cons referra clinical likely ca de Susp	ider hospital al according to circumstances, suse, and risk of terioration end RAASi ^{1,2}	Urgen re Suspen	t hospital ferral d RAASi ^{1,2}	Urg hosp refe
	Once RAASi has been titrated to maximum tolerated dose,		Maintain	RAASi ^{1,2} dose	Halve RAA	Si ^{1,2} dose	
	start an SGLT2i in eligible groups as per Figure 2, unless		Repeat K	+ within 14 days	Repeat K+	within 7 days	
3. SGLT2	contraindicated (e.g. in people living with T1DM)(#(2010)		If repeat I	K+ is:	If repeat K+	+ is:	
Inhibitors ^[18,20,21,26]	Continue the SGLI Zi until dialysis or transplant ^{10,04,00} See also the Primary Care Wask on ovtra glycaemic indications		• ≤5.5 : o	ptimise	• ≤5.5: opt	imise	
	of SGLT2is.	Well and	• 5.6-5.9	^e dose ^I maintain	• 5.6-5.9	tose maintain	Con
		<50%	RAASi ¹	² dose	RAASi ^{1,2} o	dose	hos
	If T2DM, eGFR ≥25 ml/min/1.73 m², and uACR ≥3 mg/mmol	301 1130	• 6.0-6.4 RAASil	: suspend	 6.0–6.4: s RAASi12 a 	suspend and discuss	
	(≥30 mg/g), consider adding:		initiatio	n of K+ binders	initiation	of K+ binders	
	 finerenone^{IBI}—a nonsteroidal MRA that can be added as third line to an RAASi and an SGLT2i (or second line if 		with Re	nal team ^{3,4}	with Rena	l team ^{3,4}	
	SGLT2i is inappropriate or not tolerated) if serum potassium		• ≥0.5: co hospita	l referral	 bospital 	referral	
	concentration is <5.0 mmol/l, to reduce the risk of adverse	1. Note: R	- AASis include	ACEis, ARBs, pota	sium-sparing	diuretics, and M	IRAs.
4. Specific Considerations in T2DM	semaglutide ^{III} —the FLOW study demonstrated that weekly SC semaglutide I mg reduced the risk of adverse kidney and CV outcomes in T2DM and CKD ^[20]	 Suspension NICE er See NIC with a h 	d finerenone dorses potas IE <u>TA599</u> and istory of bow	if K+ ≥5.5 mmol/l, re sium binders, as the <u>TA623</u> . Note: that I rel obstruction, majo	starting at 10 y enable RAAS (+ binders are r GI surgery, c	mg when K+ is « is use in those no contraindicated or a swallowing d	<5.0 mr ot on di d in pec lisorder
	 consider if glycaemic and weight management goals are not met and/or K+ >5.0 mmol/^{[[5,10,21]} 	Table base	d on the aut	hors' clinical experie	nce and interp	pretation of clinic	al guid
	 deprescribe any DPP4 inhibitor if initiating semaglutide^[18] 		Fig	gure 2: SGLT2i	Initiation i	n CKD	
	 do not switch to semaglutide if an existing GLP-1 RA 				uACR (mg	/mmol)	
	or dulaglutide).			<20		≥20	,
		(r_	≥60	Suggested in	T2DM	Recomme	ended
	 Opportunistically check FBC/HbA_{tc}/lipids/LBTs/weight/WtHR/BP at the same time as checking U&E and ACR, to support holistic 	33 1	45-60	Suggested in	T2DM	Recomme	ended
5 Holistic Care	interventions (see the Primary Care Hacks T2DM CVRM checklist,	GFR 1.1	20-45	Recommen	nded	Recomme	ended
J. Holistic Care	as well as the Primary Care Hacks on <u>lipid management</u> and <u>LBTs</u>)	, in l	<20	Suggeste	d ^(E)	Suggest	ted ^(E)

or because of other factors, e.g. frailty, reduced life expectancy, syncope.^[1–6,12] NICE recommends targets of 120–139 mmHg (SBP) and <90 mmHg (DBP) if ACR <70 mg/mmol, recommence aligned of 126-01 ZM milling and <80 mmilling if ACR ≥10 mg/mmilling bull /11 ACR >01 mg/mmilling bull /11 ACR >01 mg/mmilling bull /11 ACR >10 mg/ renal function after commencing an SGLT21^[3,24] Consider the patient's hydration status and adjust/reduce diuretic or anti-BP doses if high risk of hypovolaemia.^[24] As eGFR drops below 45 ml/min/1.73 m², SGLT2is' glycaemic efficacy reduces.²⁸¹ However, SGLT2is can be continued if tolerated until RRT.^[18,00,26]

[D] Relative contraindications to SGLT2i use include immunosuppression (e.g. kidney transplantation, lupus, vasculitis) and ADPKD.^[24] [E] Do not discontinue SGLT2i if renal function deteriorates (eGFR <20 ml/min/1.73 m?), as long as the SGLT2i was initiated prior to this.^[26]

ACEI=angiotensin-converting enzyme inhibitor; ACR=albumin to creatinine ratio; ADPKD=autosomal dominant polycystic kidney disease; AKI=acute kidney injury; ARB=angiotensin II receptor blocker; BP=blood pressure; KKD=chronic kidney disease; CV=cardiovascular (CVD=cardiovascular disease; DBP=distolb blood pressure; DKA=diabetic ketoacidosis; DPP4=dispetidyl peptidase 4; defR=estimated gloomerular filtration rate; BBC=full block count; Gl=gastronistensini, GUP1 Re-Jucagon-Iike peptida - Re-gotor agonist; MBA=histamice; Teceptor antagonist; Hb=haemoglobin; HbA_=haemoglobin; HbA_=haemoglobin; HSAD=nostrectiodia anti-inflammatory ding; OD=once diali); OOH=out of hours; PPI=proton=pum inhibitor; RAT=renal reglacement therapy; SADMAN5=sulforytures, ACEis, durites; metformin, ARB, NSADs, SGUT28; SBP=systolc blood pressure; SC=subcutaneous; SC=sure creatine; SGUT28=sodium=gluces oc-transporter? Inhibitor; RCS=in_achroydrate, Leknes; TIDM=hype 1 diabetes mellitus; UZE=uree and electrolytes; uACR=urine altourin to creatinine ratio; UTI=urinary tract infection; WHR=waist-to-height ratio

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w this Primary Care Hack online at medscape-uk.co/Hack-CKD			Last updated: February 2025.











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Cardiovascular, renal and metabolic conditions are all interlinked



CKD, chronic kidney disease; CVD, cardiovascular disease; DM, diabetes mellitus; T2DM, type 2 diabetes mellitus; UK, United Kingdom.

1. British Heart Foundation. UK Factsheet September 2024. Available at: <u>https://www.bhf.org.uk/what-we-do/our-research/heart-statistics</u>. Accessed October 2024; 2. Diabetes UK. Number of people living with diabetes in the UK tops 5 million for the first time. Available at: <u>https://www.diabetes.org.uk/about_us/news/number-people-living-diabetes-uk-tops-5-million-first-time</u>. Accessed October 2024; 3. Kidney Care UK. Key facts about kidneys. Available at: <u>https://www.kidneycareuk.org/news-and-campaigns/facts-and-stats/</u>. Accessed October 2024; 4. Usman MS, et al. The Interplay Between Diabetes, Cardiovascular Disease, and Kidney Disease. In: Chronic Kidney Disease and Type 2 Diabetes. American Diabetes Association 2021 Arlington (VA). Available at: <u>https://www.ncbi.nlm.nih.gov/books/NBK571718/</u>. Accessed October 2024.

Cardiovascular, renal and metabolic conditions are all interlinked



CKD, chronic kidney disease; CVD, cardiovascular disease; DM, diabetes mellitus; T2DM, type 2 diabetes mellitus; UK, United Kingdom.

1. British Heart Foundation. UK Factsheet September 2024. Available at: https://www.bhf.org.uk/what-we-do/our-research/heart-statistics. Accessed October 2024; 2. Diabetes UK. Number of people living with diabetes in the UK tops 5 million for the first time. Available at: https://www.diabetes.org.uk/about_us/news/number-people-living-diabetes-uk-tops-5-million-first-time. Accessed October 2024; 3. Kidney Care UK. Key facts about kidneys. Available at: https://www.kidneycareuk.org/news-and-campaigns/facts-and-stats/. Accessed October 2024; 4. Usman MS, et al. The Interplay Between Diabetes, Cardiovascular Disease, and Kidney Disease. In: Chronic Kidney Disease and Type 2 Diabetes. American Diabetes Association 2021 Arlington (VA). Available at: https://www.ncbi.nlm.nih.gov/books/NBK571718/. Accessed October 2024.



Chronic kidney disease, indicated by an eGFR <60 mL/min/1.73m²,* means that your patient....

- Is more likely to have ischaemic heart disease, heart failure, peripheral vascular disease or cerebrovascular disease²
- Has a higher risk of cardiovascular mortality²
- Can find it more difficult to achieve blood pressure targets³
- May be prone to ankle swelling and fluid retention⁴
- May be at increased risk of sustaining a hip fracture⁵
- Is at increased risk of hypoglycaemia⁶

^{*}Chronic kidney disease may also be diagnosed in patients with an eGFR >60 mL/min/1.73m² but who have markers of kidney damage (micro- or macroalbuminuria) 1. Wang Y, et al. *Kidney Int.* 2014; 85(5): 1192–1199; 2. Wright J and Hutchison A. *Vasc Health Risk Manag.* 2009; 5:713–722; 3. McCullough PA et al. *Curr Diab Rep.* 2011; 11(1):47–55; 4. Renal Resource Centre. Understanding Chronic Kidney Disease. Available at: <u>https://kidney.org.au/cms_uploads/docs/rrc-understanding-chronic-kidney-disease.pdf</u> (Accessed July 2020) 5. Nickolas TL et al. *J Am Soc Nephrol.* 2006; 17(11):3223–3232; 6. Moen MF et al. *Clin J Am Soc Nephrol.* 2009;4(6):1121–1127.

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*Chronic kidney dise 1. Wang Y, et al. Kid 11(1):47-55; 4. Ren (Accessed July 202





ninuria) Diab Rep. 2011; nic-kidney-disease.pdf

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ACE, angiotensin-converting enzyme; ACR, albumin to creatinine ratio; ARB, angiotensin receptor blocker; SGLT2, sodium-glucose cotransporter-2. Adapted from: NICE. Type 2 diabetes in adults: management [NG28]. Available at: <u>https://www.nice.org.uk/guidance/ng28</u>. Accessed October 2024.



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& Dr Andrew H. Frankel

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DM, diabetes mellitus; eGFR, estimated glomerular filtration rate.

1. UKKA. UK Kidney Association Clinical Practice Guideline: Sodium-Glucose Co-transporter-2 (SGLT-2) Inhibition in Adults with Kidney Disease. Available at: <u>https://guidelines.ukkidney.org/</u>. Accessed October 2024; 2. UKKA. Summary of Recommendations. Available at: <u>https://guidelines.ukkidney.org/summary-of-recommendations/</u>. Accessed October 2024.

Summary of Recommendations

Recommendations for Use in people with Type 2 DM Section 2 PEOPLE WITH TYPE 2 DM Grade We recommend initiating SGLT-2 inhibition in people with chronic kidney disease and type 2 diabetes, irrespective of primary kidney disease,* for any of the following 4 clinical scenarios: a) eGFR of 20-45 mL/min/1.73m² b) eGFR of >45 mL/min/1.73m² and a urinary albumin-to-creatinine ratio (uACR) of ≥25 mg/mmol⁺ 1A 1. c) Symptomatic heart failure, irrespective of ejection fraction d) Established coronary disease We suggest initiating SGLT-2 inhibition to modify cardiovascular risk and slow rate of kidney 2B 2. function decline in people with an eGFR >45-60 mL/min/1.73m² and a uACR of <25 mg/mmol, recognising effects on glycaemic control will be limited. We suggest clinicians consider initiating SGLT-2 inhibition in people with an eGFR below 20 2B 3. mL/min/1.73m² to slow progression of kidney disease.

Recommendations for Use in people without DM

Secti	on 3 PEOPLE WITHOUT DM	
1.	 We recommend initiating SGLT-2 inhibition in people with chronic kidney disease, irrespective of primary kidney disease,* for any of the following clinical scenarios: (a) eGFR of ≥20 mL/min/1.73m² and a urinary albumin-to-creatinine ratio (uACR) of ≥25 mg/mmol⁺ (b) Symptomatic heart failure, irrespective of ejection fraction 	1A
2.	We recommend initiating SGLT-2 inhibition to slow rate of kidney function decline in people with an eGFR of 20-45 mL/min/1.73m ² and a uACR of <25 mg/mmol ⁺ .	18
3.	We suggest clinicians consider initiating SGLT-2 inhibition in people with an eGFR below 20 mL/min/1.73m ² to slow progression of kidney disease.	2B
* exclu † urina	des people with polycystic kidney disease, type 1 diabetes, or a kidney transplant ry protein-to-creatinine ratio of 35 mg/mmol can be considered equivalent	

Author: Dr Kevin Fernar	ndo, GP Partner, North Be	rwick Health Ce	entre; Cor	tent Advisor, Medsca	pe Global and UK. Email	: kfernando@webmd.net	
		No dose adjust	tment nee	ded 🛛 😑 Dose adjustn	ent or further action recon	nmended Not recomm	ended
				CKD stage (ml/min/m			
	Stages G1 and G2 eGFR ≥60	Stage G eGFR 45	3a -59	Stage G3b eGFR 30-44	Stage G4 eGFR 15-30	Stage G5 eGFR <15	
Metformin	3 g total maximum daily dose (in 2–3 daily doses)	2 g total max daily dose (in daily doses)	imum 2–3	1 g total maximum daily dose (in 2–3 daily doses)			
Sulfonylureas		Increased risk Consider redu glipizide prefe	of hypog uting dos erred as n	lycaemia if eGFR <60 e. Gliclazide and tetabolised in the live			
Repaglinide							
Acarbose					Avoid if	CrCl <25 ml/min/1.73 m ²	
Pioglitazone	/					Avoid in those on dialysis	1
Alogliptin		Re ≤5	duce to 1 i0 ml/min	2.5 mg od if CrCl	Reduce to 6.25 mg o dialysis required	od if CrCl <30 ml/min or	
Linagliptin							
Saxagliptin		Reduce to 2.5	mg od	durante RC	Participation of the	Avoid in those on dialysis	1
Sitagliptin			Re	duce to 50 mg od	Reduce to 25 mg od		\sim
viidagiiptin	/	HOP	duce to a	o mg od if Ortil <50 r	ni/min		
Canaglifiozin	Initiate 100 mg and titrate to 300 mg if additional glycsemic improvement required	Initiate or co 100 mg only	ntinue	All SOLT2 inhibitors have negligible glucose-lowering effects once eGFR falls below 45. Consider adding an additional glucose-lowering agent if further glycaemic improvement is required Consist SOLT2 liabititors have beneficial cardio-ment effects at all strong			
Dapagliflozin	Recommended dose is	10 mg	of renal impairment and should be continued See The Medicape UK Primary Care Hack Extra-Glycaemic Indications of SGLT2 Inhibitors, for use of SGLT2 inhibitors in this context				
Empagliflozin	Initiate 10 mg and titrate to 25 mg if additional glycaemic improvement required	Initiate or continue 10 mg only		For further information, see: Diabetes Management in Chronic Kidney Disease: A Consensus, Report by the American Diabetes Association and Kidney Disease; Improving Ciabal Outcommes			
Ertugliflozin	Initiate 5 mg and titrate glycaemic improvemen initiate if eGFR <60	a to 15 mg if ad nt required. Do r	lditional not	Management of Hy sus Report by the A Association for the	perglycemia in Type 2 D American Diabetes Assor Study of Diabetes	liabetes, 2022. A Consen- clation and the European.	
Dulaslutida au	·						2
Exceptide hid			Dose esc proceed	alation should conservatively if			ſ
Exenative bio			CrCl 30-!	iD ml/min			
Exenatide qw							<
Lixisenatide od							
Semaglutide sc qw							ſ.
Semaglutide oral od		Umited exper	tence in p	atients with severe re	nal impairment eGFR <30		
Tirzepatide qw	No dose adjustment is tirzepatide in patients	required for pa with severe rena	itients wit al impairn	h renal impairment in hent and ESRD is limit	cluding ESRD. Experience ed	with the use of	
Degludec + liraglutide (Xultophy8)		Intensify gluc	ose moni	toring and dose adjus	t on an individual basis		
Glargine + lixisenatide (Suliqua®)		Intensify gluc adjust on an i	ose moni individual	toring and dose basis			
	Intensify glucose monitoring and dose adjust on an individual basis due to increased risk of hypoglycaemia						

The glucose-lowering efficacy of all SGLT2 inhibitors is dependent on renal function and is negligible when eGFR <45

If eGFR falls <45, additional glucose-lowering treatment should be considered in people living with T2D



2023 ESC Guidelines for the management of cardiovascular disease in patients with diabetes

Official ESC Guidelines slide set



Figure 1: Management of CVD in patients with type 2 diabetes: clinical approach and key recommendations



ASCVD, atherosclerotic cardiovascular disease; CKD, chronic kidney disease; CVD, cardiovascular disease; ESC, European Society of Cardiology; GLP-1 RA, glucagon-like peptide-1 receptor agonist; HF, heart failure; HFmrEF, heart failure with mildly reduced ejection fraction; HFpEF, heart failure with preserved ejection fraction; HFrEF, heart failure with reduced ejection fraction; SGLT2, sodium-glucose cotransporter-2; T2DM, type 2 diabetes mellitus. Marx N, et al. Eur Heart J 2023;44:4043-4140.

COMPREHENSIVE CARE IN PATIENTS WITH DIABETES AND CKD

Practice Point 1.1.1: Patients with diabetes and chronic kidney disease (CKD) should be treated with a comprehensive strategy to reduce risks of kidney disease progression and cardiovascular disease (Figure 1 and 2).





Three pillars approach to management of CKD associated with T2D



*If criteria met and within licence, unless unsuitable.¹ Refer to relevant SGLT2i SmPC before prescribing due to variability in licenses. †If criteria met and within licence.²

ACEi, angiotensin converting enzyme inhibitor; ARB, angiotensin receptor blocker; CKD, chronic kidney disease; eGFR, estimated glomerular filtration rate; SGLT2i, sodium-glucose cotransporter 2 inhibitor; SmPC, summary of product characteristics; T2D, type 2 diabetes; UACR, urinary albumin to creatinine ratio.

1. NICE. Type 2 diabetes in adults: management [NG28]. Available at: https://www.nice.org.uk/guidance/ng28. Accessed October 2024; 2. NICE. Finerenone for treating chronic kidney disease in type 2 diabetes [TA877]. Available at: https://www.nice.org.uk/guidance/ta877. Accessed October 2024.

WHO SHOULD BE TESTED FOR CKD?

Offer testing using eGFR, creatinine & ACR if these risk factors present:

- Diabetes
- Hypertension
- Previous AKI
- CVD
- Structural renal tract disease, recurrent renal calculi or BPH
- Multisystem disease with potential renal involvement e.g.
 SLE
- Gout
- □ FH of ESRD or hereditary kidney disease
- Incidental haematuria or proteinuria

Pitfalls & Cautions When Interpreting eGFR¹

- It is only an <u>estimate</u> and significant error is possible...
- Most likely to be inaccurate in people at extremes of body type e.g. limb amputations, malnourished or morbidly obese
- Also AKI, pregnancy, oedematous states & certain ethnic groups
- Identifying trends in eGFR is often more informative

AKI, acute kidney injury; eGFR, estimated glomerular filtration rate.

1. UKKA. Measurement of kidney function. Available at: https://ukkidney.org/health-professionals/information-resources/uk-eckd-guide/measurement-kidney-function. Accessed October 2024.

Pitfalls & Cautions When Interpreting eGFR



eGFR, estimated glomerular filtration rate.

1. UKKA. Measurement of kidney function. Available at: https://ukkidney.org/health-professionals/information-resources/uk-eckd-guide/measurement-kidney-function. Accessed October 2024; 2. Naguib R and Elkemary E. Cureus 2023;15:e35242; 3. Roncal-Jimenez C, et al. Ann Nutr Metab 2015;66:10–13.

Pitfalls & Cautions When Interpreting Urinary ACR

- Send off sample in plain urine container^{1,2}
 - <u>No</u> need for early morning sample
 - Can be stored overnight in fridge
 - ACR>3mg/mmol is clinically important proteinuria in PLWD¹
 - 2 positive samples over 3 months
- Do not use reagent strips to identify proteinuria!¹
- Why do we need 2 samples?

ACR, albumin to creatinine ratio; $\ensuremath{\mathsf{PLWD}}$, people living with diabetes.

1. Royal United Hospitals Bath NHS Foundation Trust. Screening Patients for Proteinuria. Available at: https://www.ruh.nhs.uk/pathology/documents/clinical_guidelines/PATH-

⁰⁰⁵ Screening Patients for Proteinuria%20Guideline.pdf?t=15010.9. Accessed October 2024; 2. Gloucestershire Hospitals NHS Foundation Trust. Available at: https://www.gloshospitals.nhs.uk/our-services/services-weoffer/pathology/tests-and-investigations/albumincreatinine-ratio-acr-and-proteincreatinine-ratio-pcr/. Accessed October 2024.

Causes of transient albuminuria:

Biolog	ical	Recent	Recent heavy Acu		Acute fl	uid loads
variab	oility	exerc	exercise		or di	iuresis
Orthost	Orthostatic		UTI / any febrile		Decompensated	
proteir	proteinuria		illness		heart failure	
	Menstru	Jation	Acute e in BP c glue	ele or co	evation blood ose	

BP, blood pressure; UTI, urinary tract infection.

1. UKKA. UK Kidney Association Clinical Practice Guideline: Sodium-Glucose Co-transporter-2 (SGLT-2) Inhibition in Adults with Kidney Disease. Available at:

https://ukkidney.org/sites/renal.org/files/UKKA%20guideline_SGLT2i%20in%20adults%20with%20kidney%20disease%20v1%2020.10.21.pdf. Accessed October 2024; 2. Park S, et al. Front Cardiovasc Med 2022;9:882599; 3. Haider MZ and Aslam A. Proteinuria. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing. Available at: https://www.ncbi.nlm.nih.gov/books/NBK564390/. Accessed October 2024.

Lipid-lowering

NICE National Institute for Health and Care Excellence NICE guideline Cardiovascular disease: risk assessment and reduction, including lipid modification NICE guideline Published: 14 December 2023 www.nice.org.uk/guidance/ng238 INICE 2024. All rights reserved. Subject to Notice of rights (https://www.nice.org.uk/terms-a conditions#notice-of-rights).

Atorvastatin 20mg for primary or secondary prevention of CVD for people with CKD

□ For primary prevention, aim for >40% 1 non-HDL-C

- □ For secondary prevention, aim for LDL-C ≤2mmol/L or non-HDL ≤2.6mmol/L
- Increase dose of atorvastatin if lipid targets not met
- Agree use of higher doses with renal specialist if eGFR <30ml/min/1.73m²

CKD, chronic kidney disease; CVD, cardiovascular disease; eGFR, estimated glomerular filtration rate; HDL-C, high density lipoprotein; LDL-C, low density lipoprotein. NICE. Cardiovascular disease: risk assessment and reduction, including lipid modification [NG238]. Available at: <u>https://www.nice.org.uk/guidance/ng238</u>. Accessed October 2024.

BP-lowering (adults)



- T2D & hypertension: commence ACEi or ARB irrespective of age or ethnicity²
- For people aged \geq 80 years, aim for a clinic BP <150/90mmHg²

ACEi, angiotensin converting enzyme inhibitor; ACR, albumin to creatinine ratio; ARB, angiotensin receptor blocker; BP, blood pressure; CKD, chronic kidney disease; NICE, National Institute for Health and Care Excellence; T2D, type 2 diabetes.

1. NICE. Chronic kidney disease: assessment and management [NG203]. Available at: https://www.nice.org.uk/guidance/ng203. Accessed October 2024; 2. NICE. Hypertension in adults: diagnosis and management [NG136]. Available at: https://www.nice.org.uk/guidance/ng136. Accessed October 2024.

Oral antiplatelets



Offer to adults with CKD for CVD secondary prevention

But be aware of **1** risk of bleeding

CKD, chronic kidney disease; CVD, cardiovascular disease; NICE, National Institute for Health and Care Excellence, NICE. Chronic kidney disease: assessment and management [NG203]. Available at: <u>https://www.nice.org.uk/guidance/ng203</u>. Accessed October 2024.

HbA1c targets

• Tight glycaemic control reduces risk of microvascular disease

However, no consensus on glycaemic targets

- Little evidence HbA1c <53mmol/mol reduces progression especially if ≥CKD G3
- Individualised HbA1c targets

NOTE

Anaemia in CKD may invalidate HbA1c assay

CKD, chronic kidney disease; DN, diabetic nephropathy; GLP-1, glucagon-like peptide-1; HbA1c, glycated haemoglobin; SGLT-2, sodium-glucose cotransporter 2. 1. Winocour P, et al. Br J Diabetes 2018;18:78–89; 2. Cochrane SR CD10137 2017.

HbA1c targets

• Tight glycaemic

However, no co

- Little evidence | ™ especially if ≥Ck
- Individualised H

NOTE Anaemia in CK

Table 1	Glycaemic targets in pa diabetic nephropathy-	atients with diabetes and chronic kidney disease (DN-CKD)	
	Glycaemic target	Note	1
Type 1 diabetes	48–58 mmol/mol (6.5–7.5%)	Younger patients within 10 years' duration of diabetes and variable microalbuminuria– CKD stage 2	
	58–62 mmol/mol (7.5–7.8%)	The majority of patients with proteinuria and/or CKD stages 3–4	vascular disease
	58–68 mmol/mol (7.5–8.5%)	Patients with CKD stage 5 dialysis	ts
Type 2 diabetes	48–58 mmol/mol (6.5–7.5%)	For the majority of patients who are aged <40 years, or have CKD stages 1–2 (no basis to aim for <52 mmol/mol (6.9%) unless the patient is aged <40 years and has CKD stages 1–2)	s progression
	52–58 mmol/mol (6.9–7.5%)	For those with CKD stages 3–4 this target may be appropriate with a GLP-1/SGLT-2 inhibitor- based treatment regime without insulin	
	58–68 mmol/mol (7.5–8.5 %)	For those with CKD stages 3–4 proteinuria who are on an insulin- based regime, and those with CKD stage 5 who are on dialysis	
GLP-1, gluc	cagon-like peptide-1; SGLT-2,	sodium glucose co-transporter-2.	porter 2.

CKD, chronic kidney disease; DN, diabetic nephropathy; GLP-1, 1. Winocour P, et al. Br J Diabetes 2018;18:78–89; 2. Cochrane SR CD10137 :



Medscape UK. Guidelines Primary Care Hacks: Pharmacological Management of Hyperglycaemia in People Living with Type 2 Diabetes and Chronic Kidney Disease. Available at: https://www.medscape.co.uk/view article/primary-care-hackspharmacological-managementhyperglycaemia-2022a10024hx. Accessed October 2024.

Author: Dr Kevin Fernan	do, GP Partner, North Be	rwick Health C	entre; Cor	tent Advisor, Medso	ape Glob	al and UK. Email: k	fernando@webmd.net
		No dose adjus	tment nee	ded 🛛 😑 Dose adjustr	ment or fi	urther action recomm	ended Not recomm
	Street 01 and 02		· · ·	CKD stage (ml/min/n			
	eGFR ≥60	eGFR 45	53a 5-59	eGFR 30-44		eGFR 15-30	eGFR <15
Metformin	3 g total maximum daily dose (in 2–3 daily doses)	2 g total mao daily dose (ir daily doses)	ximum n 2–3	1 g total maximum daily dose (in 2–3 daily doses)			
Sulfonylureas		Increased risk Consider red glipizide pref	k of hypog lucing dos ferred as n	lycaemia if eGFR <60 e. Gliclazide and netabolised in the live	0. er		
Repaglinide							
Acarbose						Avoid if C	rCl <25 ml/min/1.73 m²
Pioglitazone						م	woid in those on dialysis
Alogliptin		Re	educe to 1 50 ml/min	2.5 mg od if CrCl	Redia	duce to 6.25 mg od lysis required	if CrCl <30 ml/min or
Linagliptin							
Saxagliptin		Reduce to 2.5	5 mg od			Ao	oid in those on dialysis
Sitagliptin			Ra	educe to 50 mg od	Rec	fuce to 25 mg od	
Vildagliptin		R	educe to 5	i0 mg od if CrCl <50	ml/min		
Canagliflozin	titrate to 300 mg if additional glycaemic improvement required	Initiate or co 100 mg only	ntinue r	falls below 45. Cons further glycaemic in Certain SGLT2 inhit of renal impairment	sider add nprovem bitors hav	ing an additional gl ent is required re beneficial cardio- uld be continued	renal effects at all stage
Dapagliflozin	Recommended dose is	10 mg		See The Medscape Indications of SGU context	e UK Prir T2 Inhibi	nary Care Hack, Ex tors, for use of SGL	tra-Glycaemic T2 inhibitors in this
Empagliflozin	Initiate 10 mg and titrate to 25 mg if additional glycaemic improvement required	Initiate or co 10 mg only	intinue	For further informat Diabetes Manager Report by the Ami Improving Global I	tion, see: ment in (erican Di Outcomi	Chronic Kidney Dise abetes Association	ease: A Consensus and Kidney Disease:
Ertugliflozin	Initiate 5 mg and titrate glycaemic improvemen initiate if eGFR <60	e to 15 mg if ac nt required. Do	dditional not	Management of H sus Report by the Association for the	vperglyc America s Study o	emia in Type 2 Dia n Diabetes Associa of Diabetes	betes, 2022, A Consen- tion and the European
Dulaciutida cre	(
Exenatide bid			Dose esc proceed CrCl 30-3	alation should conservatively if 50 ml/min			
Exenatide gw							
Liraglutide od							
Lixisenatide od							
õemaglutide sc qw		Limited area	denne is s	atients with an	and in	irmant aGEP c20	
Semaglutide oral od		onneo expe	mence in p	audius with severe R	and impl	annen: eork <d0< td=""><td></td></d0<>	
Tirzepatide qw	No dose adjustment is tirzepatide in patients	required for pa with severe ren	atients wit al impairn	h renal impairment in hent and ESRD is limi	cluding I ted	ESRD. Experience w	ith the use of
Degludec + iraglutide (Xultophy8)		Intensify glu	cose moni	toring and dose adju	st on an	individual basis	
Glargine + lixisenatide (Suliqua®)		Intensify glue adjust on an	cose moni individual	toring and dose basis			
All insulins		Intensify glue	cose moni	toring and dose adju	st on an	individual basis due	to increased

Last updated: November 2022

When to consider referral?

- □ 5-year risk of needing RRT >5% measured using 4-variable KFRE
- □ Sustained I in eGFR ≥25% and change in eGFR category
- □ Sustained \downarrow in eGFR ≥15 within 12 months
- □ ACR ≥70mg/mmol unless known to be caused by diabetes & appropriately treated
- □ ACR >30mg/mmol (A3) with haematuria
- Poorly controlled BP despite 4 agents
- □ Suspected renal artery stenosis
- □ Known or suspected rare or genetic causes of CKD

□ RRT required within lifetime?

ACR, albumin to creatinine ratio; BP, blood pressure; CKD, chronic kidney disease; eGFR, estimated glomerular filtration rate; KFRE, Kidney Failure Risk Equation; RRT, renal replacement therapy. 1. NICE. Chronic kidney disease: assessment and management [NG203]. Available at: <u>https://www.nice.org.uk/guidance/ng203</u>. Accessed October 2024; 2. Calculate by QxMD. Kidney Failure Risk Equation (4 Variable). Available at: <u>https://qxmd.com/calculate/calculator_308/kidney-failure-risk-equation-4-variable</u>. Accessed October 2024. When Calculate **5**-year 🛛 Sustai 🛛 Sustai □ ACR ≥ treated □ ACR > Poorly □ Know

RRT re

Calculator

About

Results Save Copy Results Kidney Failure Risk Equation (4 Variable) \$ Risk of progression to kidney failure requiring dialysis or transplantation Estimate risk of progression to end-stage renal disease in CKD patients using age, sex, eGFR and proteinuria with KFRE **Over 2-Years:** 0% Questions 1. Sex? Male Over 5-Years: 2. Age? 45 Years 0% 3. eGFR? 90 mL/min/1.73m² Urine Albumin Creatinine Ratio? (Note units carefully) 0 mg/mmol 4. 5. Patient location? Non-North America ely About The kidney failure risk equations were developed in patients with CKD stages G3-G5 referred to nephrologists in Canada, and have now been validated in more than 700,000 individuals spanning 30 + countries worldwide. The four and eight variable equations accurately predict the 2 and 5 year probability of treated kidney failure (dialysis or transplantation) for a potential patient with CKD Stage 3 to 5. Predicted risks may differ from observed risks in clinical populations with lower and higher observed risks than the study population, and a calibration factor for non-North American cohorts has been added Determining the probability of kidney failure may be useful for patient and provider communication, triage and management of nephrology referrals and timing of dialysis access placement and living related kidney transplant. Prospective trials evaluating the utility of this instrument for clinical decision making are in progress. References Tangri N, Stevens LA, Griffith J, et al. **Try 8-Variable Kidney Failure Risk Equation** A predictive model for progression of chronic kidney disease to kidney failure. Estimate risk of progression to end-stage renal disease in CKD patients with more precision using 8 JAMA: the Journal of the American Medical Association 2011 April 20, 305 (15): 1553-9 variables. Tangri N, Grams ME, Levey AS et al. Multinational Assessment of Accuracy of Equations for Predicting Risk of Kidney Failure: A Get IT ON Google Play App Store Download the app for offline access Meta-analysis. 5 Copy Results 5/5 completed

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ACR, albumin to creatinine ratio; BP, blood pressure; CKD, chronic kidney disease; eGFR, estimated glomerular filtration rate; KFRE, Kidney Failure Risk Equation; RRT, renal replacement therapy. 1. NICE. Chronic kidney disease: assessment and management [NG203]. Available at: <u>https://www.nice.org.uk/guidance/ng203</u>. Accessed October 2024; 2. Calculate by QxMD. Kidney Failure Risk Equation (4 Variable). Available at: <u>https://qxmd.com/calculate/calculator_308/kidney-failure-risk-equation-4-variable</u>. Accessed October 2024.

References

2. RAASis and Blood Pressure^[3-5,12,18-25]

• Aim for SBP <130 mmHg, or as low as reasonably achievable^[A]

- Follow <u>NG136^[19]</u> and <u>NG203^[3]</u> when choosing medications
- When RAASi is first line, choose an **ARB**^[B]
- Independent of BP:[B]
 - if uACR >30 mg/mmol: start ARB and titrate to maximum tolerated dose
 - if uACR >3 mg/mmol in people with diabetes: start ARB and titrate to maximum tolerated dose
 - additional agents may be required
- If eGFR <60 ml/min/1.73 m² when starting or increasing RAASi, check U&E within **28 days**; if eGFR ≥60 ml/min/1.73 m², there is no need to routinely recheck U&E
 - if K+ ≥5.5 mmol/l or sCr rise ≥50% encountered, follow
 Figure 1; otherwise, continue to optimise RAASi dose.

Figure 1: Managing Acute Changes in Kidney Function Related to RAASis ^[3,22–25]					
	Serui	m K+ (mmol/l)			
	5.5–6.1	6.2–6.4	≥6.5		
Unwell and/or ≥50% sCr rise	Consider hospital referral according to clinical circumstances, likely cause, and risk of deterioration Suspend RAASi ^{1,2}	Urgent hospital referral Suspend RAASi ^{1,2}	Urgent hospital referral		
Well and <50%	Maintain RAASi ^{1,2} dose Repeat K+ within 14 days If repeat K+ is: • ≤5.5: optimise RAASi ^{1,2} dose • 5.6–5.9: maintain RAASi ^{1,2} dose	Halve RAASi ^{1,2} dose Repeat K+ within 7 days If repeat K+ is: • ≤5.5: optimise RAASi ^{1,2} dose • 5.6–5.9: maintain RAASi ^{1,2} dose	Conside hospita		
sCr rise	 6.0-6.4: suspend RAASi^{1,2} and discuss initiation of K+ binders with Renal team^{3,4} ≥6.5: consider hospital referral 	 6.0-6.4: suspend RAASi^{1,2} and discuss initiation of K+ binders with Renal team^{3,4} ≥6.5: consider hospital referral 	referral		

- 3. NICE endorses potassium binders, as they enable RAASi use in those not on dialysis.
- 4. See NICE <u>TA599</u> and <u>TA623</u>. Note: that K+ binders are contraindicated in people with a history of bowel obstruction, major GI surgery, or a swallowing disorder.

Table based on the authors' clinical experience and interpretation of clinical guidance.



Disease in Primary Care

Medscape ***** UK X Guidelines

Primary Care Hacks

Authors: Dr Eimear Darcy, GP Partner, Grange Family Practice Omagh; Dr William Hinchliffe, Consultant in Renal and General Medicine, South Tyneside and Sunderland Foundation NHS Trust; Dr Deepika Manoharan, Registrar in Renal and General Medicine, South Tyneside and Sunderland Foundation NHS Trust; Dr Kevin Fernando, GP Partner, North Berwick Health Centre and Content Advisor, Medscape Global and UK (email: kfernando@webmd.net)

	For primary prevention, start atorvastatin 20 mg OD	Lifest	yle and	Dietary Mo	dificatio	n	
1. Lipids ^[3–5,15–18]	 For secondary prevention, do not stop or attenuate dose if eGFR <30 m//min/1.73 m² Offer aspirin for secondary prevention of CVD See also the <u>Primary Care Hack on lipid management</u>. 	 Encours Smoking Advise (ideally 	age weigh ng cessatio on salt re <2 g of so	nt loss and on ^[3-6,8] striction odium per	with incre and prog • Avoid NS • Promote	ased risk of n ression of CKI AIDs ⁽³⁻⁵⁾ exercise of at	ephriti D ^(13,14) t least
	• Aim for SBP <130 mmHg , or as low as reasonably achievable ^(A) • Follow <u>NG136⁽¹⁹⁾ and NG203^[0] when choosing medications</u>	day, ec sodium • If gastr consid	uating to chloride) ^[] ic protection er H_RAs c	<5 g of 15.8,121 on is required, over PPIs, as	 150 minu Kidney Ca a helpful leaflet to 	tes per week ¹² are UK has pro patient inform support peop	aduce
	When RAASi is first line, choose an ARB ^[8] Independent of BP: ⁽⁸⁾ if uACR >30 mo/mmol: start ARB and titrate to	PPI use	has been	associated	with CKD		
2. RAASis	maximum tolerated dose	riy	ure 1. ivia	Related to F	AASis ^{[3,22-3}		cuon
and Blood	start ARB and titrate to maximum tolerated dose			Seru	m K+ (mmo	I/I)	
Pressure	additional agents may be required			5.5–6.1	6.3	2-6.4	≥6.
	 If GURK <60 m/mm/1, 1.3 m² when starting or increasing KAASI, check U&E whith 28 days; If GCFR >60 m/min/1.73 m², there is no need to routinely recheck U&E If K+ 25.5 mmol/l or sCr rise 250% encountered, follow Figure 1; otherwise, continue to optimise RAASi dose. 	Unwell and/or ≥50% sCr rise	Cons referra clinical likely ca de Susp	ider hospital al according to circumstances, suse, and risk of terioration end RAASi ^{1,2}	Urgen re Suspen	t hospital ferral d RAASi ^{1,2}	Urg hosp refe
	Once RAASi has been titrated to maximum tolerated dose,		Maintain	RAASi ^{1,2} dose	Halve RAA	Si ^{1,2} dose	
	start an SGLT2i in eligible groups as per Figure 2, unless		Repeat K	+ within 14 days	Repeat K+	within 7 days	
3. SGLT2	contraindicated (e.g. in people living with T1DM)(#(2010)		If repeat I	K+ is:	If repeat K+	+ is:	
Inhibitors ^[18,20,21,26]	Continue the SGLI Zi until dialysis or transplant ^{10,04,00} See also the Primary Care Wask on ovtra glycaemic indications		• ≤5.5 : o	ptimise	• ≤5.5: opt	imise	
	of SGLT2is.	Well and	• 5.6-5.9	^e dose ^I maintain	• 5.6-5.9	tose maintain	Con
		<50%	RAASi ¹	² dose	RAASi ^{1,2} o	dose	hos
	If T2DM, eGFR ≥25 ml/min/1.73 m², and uACR ≥3 mg/mmol	301 1130	• 6.0-6.4 RAASil	: suspend	 6.0–6.4: s RAASi12 a 	suspend and discuss	
	(≥30 mg/g), consider adding:		initiatio	n of K+ binders	initiation	of K+ binders	
	 finerenone^{IBI}—a nonsteroidal MRA that can be added as third line to an RAASi and an SGLT2i (or second line if 		with Re	nal team ^{3,4}	with Rena	l team ^{3,4}	
	SGLT2i is inappropriate or not tolerated) if serum potassium		• ≥0.5: co hospita	l referral	 bospital 	referral	
	concentration is <5.0 mmol/l, to reduce the risk of adverse	1. Note: R	- AASis include	ACEis, ARBs, pota	sium-sparing	diuretics, and M	IRAs.
4. Specific Considerations in T2DM	semaglutide ^{III} —the FLOW study demonstrated that weekly SC semaglutide I mg reduced the risk of adverse kidney and CV outcomes in T2DM and CKD ^[20]	 Suspension NICE er See NIC with a h 	d finerenone dorses potas IE <u>TA599</u> and istory of bow	if K+ ≥5.5 mmol/l, re sium binders, as the <u>TA623</u> . Note: that I rel obstruction, majo	starting at 10 y enable RAAS (+ binders are r GI surgery, c	mg when K+ is « is use in those no contraindicated or a swallowing d	<5.0 mr ot on di d in pec lisorder
	 consider if glycaemic and weight management goals are not met and/or K+ >5.0 mmol/^{[[5,10,21]} 	Table base	d on the aut	hors' clinical experie	nce and interp	pretation of clinic	al guid
	 deprescribe any DPP4 inhibitor if initiating semaglutide^[18] 		Fig	gure 2: SGLT2i	Initiation i	n CKD	
	 do not switch to semaglutide if an existing GLP-1 RA 				uACR (mg	/mmol)	
	or dulaglutide).			<20		≥20	,
		(r_	≥60	Suggested in	T2DM	Recomme	ended
	 Opportunistically check FBC/HbA_{tc}/lipids/LBTs/weight/WtHR/BP at the same time as checking U&E and ACR, to support holistic 	33 1	45-60	Suggested in	T2DM	Recomme	ended
5 Holistic Care	interventions (see the Primary Care Hacks T2DM CVRM checklist,	GFR 1.1	20-45	Recommen	nded	Recomme	ended
J. Holistic Care	as well as the Primary Care Hacks on <u>lipid management</u> and <u>LBTs</u>)	, in l	<20	Suggeste	d ^(E)	Suggest	ted ^(E)

or because of other factors, e.g. frailty, reduced life expectancy, syncope.^[1–6,12] NICE recommends targets of 120–139 mmHg (SBP) and <90 mmHg (DBP) if ACR <70 mg/mmol, recommence aligned of 126-01 ZM milling and <80 mmilling if ACR ≥10 mg/mmilling bull /11 ACR >01 mg/mmilling bull /11 ACR >01 mg/mmilling bull /11 ACR >10 mg/ renal function after commencing an SGLT21^[3,24] Consider the patient's hydration status and adjust/reduce diuretic or anti-BP doses if high risk of hypovolaemia.^[24] As eGFR drops below 45 ml/min/1.73 m², SGLT2is' glycaemic efficacy reduces.²⁸¹ However, SGLT2is can be continued if tolerated until RRT.^[18,00,26]

[D] Relative contraindications to SGLT2i use include immunosuppression (e.g. kidney transplantation, lupus, vasculitis) and ADPKD.^[24] [E] Do not discontinue SGLT2i if renal function deteriorates (eGFR <20 ml/min/1.73 m?), as long as the SGLT2i was initiated prior to this.^[26]

ACEI=angiotensin-converting enzyme inhibitor; ACR=albumin to creatinine ratio; ADPKD=autosomal dominant polycystic kidney disease; AKI=acute kidney injury; ARB=angiotensin II receptor blocker; BP=blood pressure; KKD=chronic kidney disease; CV=cardiovascular (CVD=cardiovascular disease; DBP=distolb blood pressure; DKA=diabetic ketoacidosis; DPP4=dispetidyl peptidase 4; defR=estimated gloomerular filtration rate; BBC=full block count; Gl=gastronistensini, GUP1 Re-Jucagon-Iike peptida - Re-gotor agonist; MBA=histamice; Teceptor antagonist; Hb=haemoglobin; HbA_=haemoglobin; HbA_=haemoglobin; HSAD=nostrectiodia anti-inflammatory ding; OD=once diali); OOH=out of hours; PPI=proton=pum inhibitor; RAT=renal reglacement therapy; SADMAN5=sulforytures, ACEis, durites; metformin, ARB, NSADs, SGUT28; SBP=systolc blood pressure; SC=subcutaneous; SC=sure creatine; SGUT28=sodium=gluces oc-transporter? Inhibitor; RCS=in_achroydrate, Leknes; TIDM=hype 1 diabetes mellitus; UZE=uree and electrolytes; uACR=urine altourin to creatinine ratio; UTI=urinary tract infection; WHR=waist-to-height ratio

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ACR=albumin to creatinine ratio; AKI=acute kidney injury; BP=blood pressure; CKD=chronic kidney disease; CNI=calcineurin inhibitor; CVD=cardiovascular disease; CVE=cardiovascular event; eGR=estimated glomerular filtration rate; EPO=erythropoietin; ESRD=end=stage renal disease; GFR=glomerular filtration rate; KFRE=Kidney Failure Risk Equation; NSAID=nonsteroidal anti-inflammatory drug; NVH=non-visible haematuria; PKD=polycystic kidney disease; QoL=quality of life; RAG=red, amber; green; sCr=serum creatinine; uACR=urine albumin to creatinine ratio; UTI=urinary tract infection

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Thank-you for listening. Any questions?

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