Interventions for Chronic Kidney Disease in Primary Care

Medscape # UK X Guidelines Primary Care Hacks

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1. Lipids^[3–5,15–18]

- For primary prevention, start atorvastatin 20 mg OD
- For secondary prevention, do not stop or attenuate dose if eGFR <30 ml/min/1.73 m²
- Offer aspirin for secondary prevention of CVD
- See also the Primary Care Hack on lipid management.

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

- Aim for SBP <130 mmHg, or as low as reasonably achievable^[A]
- Follow NG136^[19] and NG203^[3] 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
 - $\circ\,$ if uACR >3 mg/mmol in people with diabetes: start ARB and titrate to maximum tolerated dose
 - · additional agents may be required
- \bullet 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.

3. SGLT2

- Once RAASi has been titrated to maximum tolerated dose, start an SGLT2i in eligible groups as per Figure 2, unless contraindicated (e.g. in people living with T1DM)[B],[C],[D]
- Inhibitors [18,20,21,26] Continue the SGLT2i until dialysis or transplant [18,20,26]
 - See also the Primary Care Hack on extra-glycaemic indications

4. Specific **Considerations** in T2DM

If T2DM, eGFR \geq 25 ml/min/1.73 m², and uACR \geq 3 mg/mmol (≥30 mg/g), consider adding:

- finerenone^[B]—a nonsteroidal MRA that can be added as third line to an RAASi and an SGLT2i (or second line if SGLT2i is inappropriate or not tolerated) if serum potassium concentration is <5.0 mmol/l, to reduce the risk of adverse kidney and CV outcomes[6,8,18,20,27,28]—see bit.ly/3v8PX4f
- **semaglutide**^[B]—the FLOW study demonstrated that weekly SC semaglutide 1 mg reduced the risk of adverse kidney and CV outcomes in T2DM and CKD[29]
 - consider if glycaemic and weight management goals are not met and/or K+ >5.0 mmol/|^[5,18,21]
 - deprescribe any DPP4 inhibitor if initiating semaglutide^[18]
 - do not switch to semaglutide if an existing GLP-1 RA with proven CV benefit is achieving goals (e.g. liraglutide or dulaglutide).

5. Holistic Care

- \bullet Opportunistically check FBC/HbA $_{\mbox{\tiny t}}$ /lipids/LBTs/weight/WtHR/BP at the same time as checking U&E and ACR, to support holistic interventions (see the Primary Care Hacks T2DM CVRM checklist, as well as the Primary Care Hacks on lipid management
- Offer vaccinations according to the <u>Green Book</u>. [30,31]

Lifestyle and Dietary Modification

- Encourage weight loss and smoking cessation[3-
- Advise on salt restriction (ideally <2 g of sodium per day, equating to <5 g of sodium chloride)^[3-5,8,12]
- If gastric protection is required, consider H, RAs over PPIs, as PPI use has been associated
- with increased risk of nephritis and progression of CKD^[13,14]
- Avoid NSAIDs[3-5]
- Promote **exercise** of at least 150 minutes per week^{[3-6,}
- Kidney Care UK has produced a helpful patient information leaflet to support people living with CKD.

Figure 1: Managing Acute Changes in Kidney Function Related to RAASis

	Serum 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% sCr rise	Maintain RAASi ^{1,2} dose Repeat K+ within 14 days If repeat K+ is:	Halve RAASi¹.² dose Repeat K+ within 7 days If repeat K+ is:	Consider hospital referral

- 1. Note: RAASis include ACEis, ARBs, potassium-sparing diuretics, and MRAs.
- 2. Suspend finerenone if $K+ \ge 5.5$ mmol/l, restarting at 10 mg when K+ is <5.0 mmol/l. 3. NICE endorses potassium binders, as they enable RAASi use in those not on dialysis
- See NICE TA599 and TA623. 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.

Figure 2: SGLT2i Initiation in CKD			
	uACR (mg/mmol)		
	<20	≥20	
≥60	Suggested in T2DM	Recommended	
45–60	Suggested in T2DM	Recommended	
20–45	Recommended	Recommended	
<20	Suggested ^[E]	Suggested ^[E]	
Dialysis	Not recommended ^[E]		

Footnotes

- [A] BP target can be relaxed and individualised if the patient cannot tolerate SBP <130 mmHg or because of other factors, e.g. frailty, reduced life expectancy, syncope. [3-5,12] NICE recommends targets of 120–139 mmHg (SBP) and <90 mmHg (DBP) if ACR <70 mg/mmol, and targets of 120–129 mmHg and <80 mmHg if ACR ≥70 mg/mmol. [3] However, in various studies, intense SBP control has not resulted in compromise. [3.6,12]

 [B] Provide advice on SICK day rules and review the SADMANS mnemonic (to include finerenone and GLP-1 RAs). See bit.ly/4bRbejE. [18,32]

 [C] Counsel patient on the main side effects of SGLT2i use, including risk of UTIs, mycotic genital infections, Fournier's gangrene, DKA, foot disease, and dehydration. [26,33] Provide

- SICK day guidance, including a leaflet (bit.ly/4bRbejF). [26,32] DO NOT routinely check renal function after commencing an SGLT2^(1,1,2,2) Consider the patient's hydration status and adjust/reduce diuretic or anti-BP doses if high risk of hypovolaemia.⁽²⁶⁾ As eGFR drops below 45 ml/min/1.73 m², SGLT2is' glycaemic efficacy reduces. [28] However, SGLT2is can be continued if tolerated until RRT. [18,20,26]
- [D] Relative contraindications to SGLT2i use include immunosuppression (e.g. kidney transplantation, lupus, vasculitis) and ADPKD. [28]
 [E] Do not discontinue SGLT2i if renal function deteriorates (eGFR <20 ml/min/1.73 m²), 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; CKD=chronic kidney disease; CV=cardiovascular; CVD=cardiovascular disease; DBP=diastolic blood pressure; DKA=diabetic ketoacidosis; DPP4=dipeptidyl peptidase-4; eGFR=estimated glomerular filtration rate; FBC=full blood count; GI=gastrointestinal; GLP-1 RA=glucagon-like peptide-1 receptor agonist; H₂RA=histamine-2 receptor antagonist; Hb=haemoglobin; HbA_{1c}=haemoglobin A1c; HF=heart failure; KDIGO=Kidney Disease: Improving Global Outcomes; K+=potassium; LBT=liver blood test; MRA=mineralocorticoid receptor antagonist; NG=NICE Guideline; NSAID=nonsteroidal anti-inflammatory drug; OD=once daily; OOH=out of hours; PPI=proton-pump inhibitor; RAASi=renin-angiotensin-aldosterone system inhibitor; RRT=renal replacement therapy; SADMANS=sulfonylureas, ACEis, diuretics, metformin, ARBs, NSAIDs, SGIT2is; SBP=systolic blood pressure; SC=subcutaneous; sCr=serum creatinine; SGIT2i=sodium-glucose co-transporter-2 inhibitor; SICK=sugar, insulin, carboylorate, ketones; T1DM=type 1 diabetes mellitus; T2DM=type 2 diabetes mellitus; U&E=urea and electrolytes; UACP=uring albumin to creatinine ratio. UTII-uring treat infection. uACR=urine albumin to creatinine ratio; UTI=urinary tract infection; WtHR=waist-to-height ratio