

Chronic Kidney Disease

Definition of CKD

- Duration of >3months plus either:
 - Kidney Damage evidenced by:
 - Albuminuria: UACR > 30mg/g
 - Structural abnormality: tumors, scarring, cysts
 - Pathological abnormality (evidence of pathology based on biopsy)
 - Decreased Kidney Function: eGFR < 60mL/min/1.73 m²

Common Causes of CKD

- Diabetes
- Hypertension
- Glomerulonephritis
- Interstitial nephritis
- Polycystic kidney disease or other inherited kidney diseases
- Prolonged obstruction of the urinary tract, from conditions such as enlarged prostate, kidney stones and some cancers
- Vesicoureteral reflux
- Recurrent kidney infection (pyelonephritis)

When to Refer

- eGFR <30 mL/min/1.73 m².
- Rapid progression (>5 mL/min/1.73 m² decline in eGFR in 12 months).
- Persistent albuminuria (ACR >300 mg/g) despite therapy.
- Refractory hypertension or hyperkalemia despite management.
- <https://www.kidneyfailure.com/>

| GFR (mL/min/1.73 m ²) | A1 <30 mg/g (Normal/Mild) | A2 30–300 mg/g (Moderate) | A3 >300 mg/g (Severe) |
|-----------------------------------|---------------------------|---------------------------|-----------------------|
| G1 (≥90) | ● Low risk | ● Moderate risk | ● High risk |
| G2 (60–89) | ● Low risk | ● Moderate risk | ● High risk |
| G3a (45–59) | ● Moderate risk | ● High risk | ● Very high risk |
| G3b (30–44) | ● High risk | ● Very high risk | ● Very high risk |
| G4 (15–29) | ● Very high risk | ● Very high risk | ● Kidney failure risk |
| G5 (<15) | ● Kidney failure risk | ● Kidney failure risk | ● Kidney failure risk |

CKD: Holistic Approach

- Lifestyle:** Healthy diet, Exercise, Weight management, Avoid tobacco
- First Line Rx** for most patients: SGLT2i⁺⁺ + SBP goal < 120* (at max tolerated ACE/ARB⁺ dose); statin: moderate to high dose
- Targeted Rx based on complications**
 - DM
 - May consider GLP-1
 - ns-MRA⁺⁺ if needed for BP goal or if albuminuria persists
 - HTN
 - After ACE/ARB⁺ then CCB and/or diuretic followed by MRA if eGFR ≥ 45
 - ASCVD high risk
 - Antiplatelet
 - Consider ezetimibe, PCSK9i (see CAD protocol for full list)

⁺ns-MRA: non-steroidal Mineralocorticoid: finerenone (hold finerenone if Potassium rises over 5.5mmol/l)

⁺ Empagliflozin can be started down to an eGFR ≥ 20 ml/min (but once initiated, it is reasonable to continue even if eGFR falls below 20 ml/min)

^{*} Expect <30% drop in eGFR after starting ACE/ARB, finerenone, or SGLT2i in the short term. If > 30% then need to hold & evaluate cause

Cystatin C vs Creatinine for eGFR

| Feature | Creatinine-Based eGFR | Cystatin C-Based eGFR |
|---------------------------|---|---|
| Source | Muscle metabolism | Produced by all nucleated cells |
| Affected by Muscle Mass? | Yes (higher in people with more muscle mass) | No (more stable across body types) |
| Affected by Diet? | Yes (high-protein diet, creatine supplements affect levels) | No (not influenced by diet) |
| Affected by Inflammation? | No | Yes (elevated in inflammatory states, obesity, thyroid dysfunction) |
| Cost & Availability | Inexpensive, widely available | More expensive, less commonly tested |
| Guideline Preference? | Default for most eGFR equations (e.g., CKD-EPI) | KDIGO recommends combined Creatinine + Cystatin C for better accuracy |

| Medication | T2D + CKD + Albuminuria | CKD + Albuminuria (No DM) | CKD Alone |
|----------------------------|---|--|--|
| RASi (ACEi/ARB) | First-line | First-line | May be used if hypertension or proteinuria |
| SGLT2i | Strongly recommended | Recommended if albuminuria ≥200 mg/g | May be considered if eGFR ≥20 |
| MRA (nsMRA or sMRA) | Add if albuminuria persists despite RASi + SGLT2i; prefer nsMRA | Not recommended for non-diabetic CKD (nsMRA); sMRA can be used for resistant HTN | Not routinely used |
| GLP-1 RA | Consider if additional glycemic or cardiovascular benefit is needed | Not recommended yet | Not recommended yet |

Common Changes in Chronic Kidney Disease (CKD): Causes and Treatments

| Change | Cause | Treatment |
|---|--|--|
| Anemia | Decreased erythropoietin (EPO) production by kidneys | Erythropoiesis-stimulating agents (ESAs), iron supplementation |
| Hyperparathyroidism | Reduced kidney function → phosphate retention, impaired vitamin D activation, hypocalcemia | Phosphate binders, vitamin D supplements, calcimimetics |
| Hypocalcemia | Impaired activation of vitamin D and phosphate retention | Calcium supplements, active vitamin D (calcitriol) |
| Hyperphosphatemia | Reduced kidney function → phosphate excretion impairment | Phosphate binders (e.g., calcium carbonate), dietary phosphate restriction |
| Fluid Retention (Edema) | Reduced sodium and water excretion due to impaired kidney function | Diuretics (e.g., furosemide), fluid restriction, sodium reduction |
| Hypertension | Sodium and water retention, impaired renin-angiotensin-aldosterone system regulation | Antihypertensives (ACE inhibitors, ARBs, diuretics) |
| Metabolic Acidosis | Reduced excretion of hydrogen ions, impaired bicarbonate reabsorption | Bicarbonate supplementation (e.g., sodium bicarbonate if HCO ₃ <22) |
| Proteinuria | Glomerular damage allows proteins to leak into urine | ACE inhibitors, ARBs, dietary protein restriction |
| Hyperkalemia | Reduced potassium excretion due to impaired kidney function | Potassium binders, dietary potassium restriction, diuretics |
| Decreased Vitamin D Activation | Impaired conversion of vitamin D to its active form | Active vitamin D (calcitriol), calcium supplementation |
| Uremia (Azotemia) | Accumulation of nitrogenous waste products due to reduced GFR | Dialysis (in advanced cases), dietary protein restriction |
| Dyslipidemia | Altered lipid metabolism in CKD | Statins, lifestyle modifications (diet, exercise) |
| Bone Mineral Disorder (Renal Osteodystrophy) | Imbalance of calcium, phosphate, and PTH, vitamin D deficiency | Phosphate binders, calcium supplements, vitamin D, PTH-lowering agents (e.g., calcimimetics) |
| Hyperuricemia (High Uric Acid) | Impaired kidney function → reduced uric acid excretion, leading to elevated levels | Uric acid-lowering therapy (e.g., allopurinol), dietary modifications (reduce purines)* * Treatment indicated only if symptomatic |

| eGFR | Type of Care | Risk of Kidney Failure |
|-------|--|------------------------|
| ≥60 | Primary Care | ≤ 3-5% |
| 30-60 | Transition from primary care to nephrology | ≥ 3-5% |
| <30 | Transition from nephrology to interprofessional care | ≥ 10% |
| <20 | Access and transplant planning | ≥ 40% |

Vaccines

- All routine vaccines plus pneumococcal vaccine

CKD Cautions

- Avoid NSAIDs
- Polypharmacy risk, avoid nephrotoxic agents
- Restrict sodium, potassium, phosphate, protein in later stage CKD (prefer plant based proteins)