

Chronic kidney disease

4th years

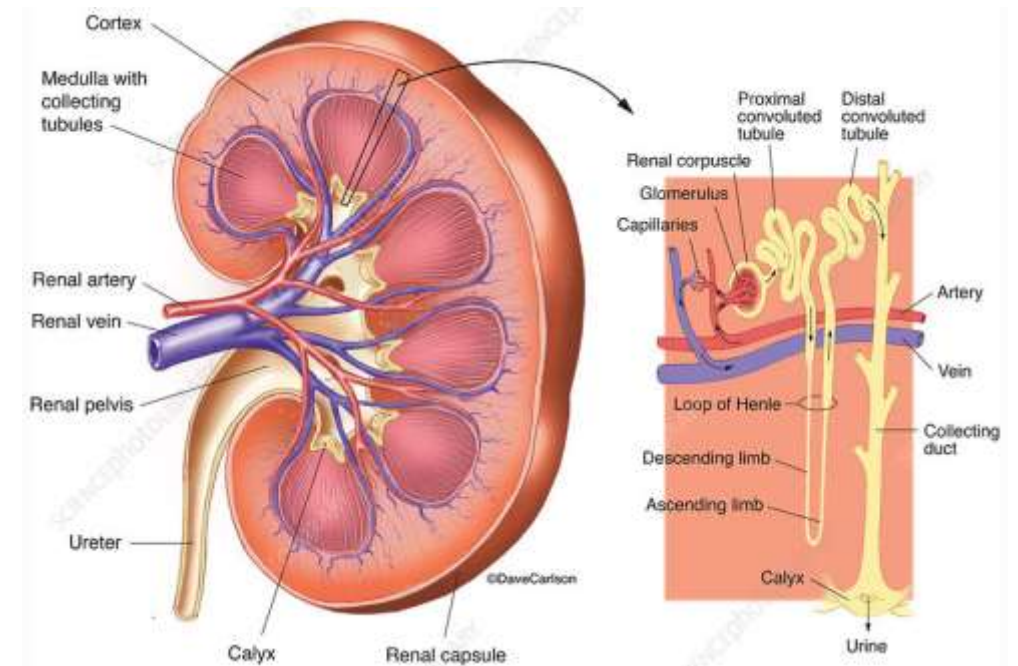
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Outline

- Basics about the kidney
- Kidney functions
- GFR (estimation versus measurement)
- Pathophysiology of kidney disease
- CKD definition/ stages
- Epidemiology/Jordan
- Manifestation
- Complication
- Management (headlines)

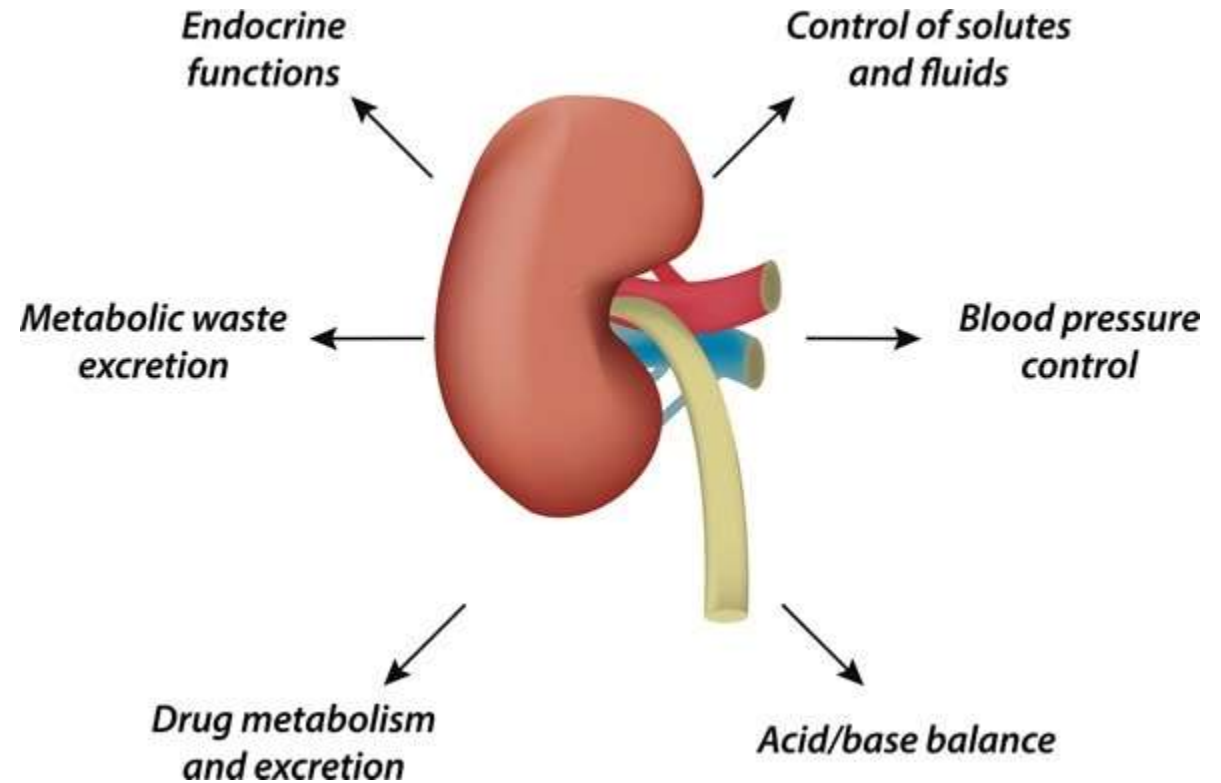
Kidney

- Kidney 150 gm (heart 300 mg), 12 cm length (left kidney larger than right)
- 1.3 million nephrons each kidney (fixed number, completed early in life)
- 2 types of nephrons
- Cortical 85%: short loop of Henle
- Juxtamedullary: long loop of Henle with Vasa recta, participate in concentration of urine



Kidney functions

- Regulation of fluid, pH, BP (short term = RAAS or long term = blood volume)
- Homeostasis of electrolytes
- Endocrine (3 hormones RAAS, activation of Vitamin D, Erythropoietin 85%) Paracrine (PG E2, NO)
- Clearance: Excretion (endogenous or exogenous waste)
- Gluconeogenesis 10% (fasting)



GFR (estimation versus measurement)

- Constant production ✓
- Freely filtered at the glomerulus ✓
- No tubular secretion or resorption
 - Some tubular secretion X
- No extra-renal metabolism ✓
- No extra-renal loss
 - Some GIT loss X
- Loss of creatinine through avenues other than glomerular filtration means Creatinine Clearance is slightly higher than the GFR

Other filtration marker (exogenous)

BOX 1

Examples of Common Glomerular Filtration Markers

Radionuclides

- Inulin labeled with carbon 14 (^{14}C -inulin)
- Technetium Tc 99m diethylenetriaminepentaacetic acid ($^{99\text{m}}\text{Tc}$ -DTPA)
- Chromium Cr 51 ethylenediaminetetraacetic acid (^{51}Cr -EDTA)

Nonradionuclides

- Iodinated
 - Ionic: Iothalamate
 - Nonionic: Iohexol, Iopamidol
- Noniodinated
 - Inulin
 - Creatinine (endogenous or exogenously administered)

CKD definition

- Progressive irreversible loss of kidney functions
- Before 2002 (KDOQI): chronic renal failure
- Structural or functional abnormalities of the kidneys for >3 months, with or without decreased GFR < 60.
- NICE – UK 2008 : stage 3 a and b
- KDIGO 2012: A1-A3

Pathophysiology

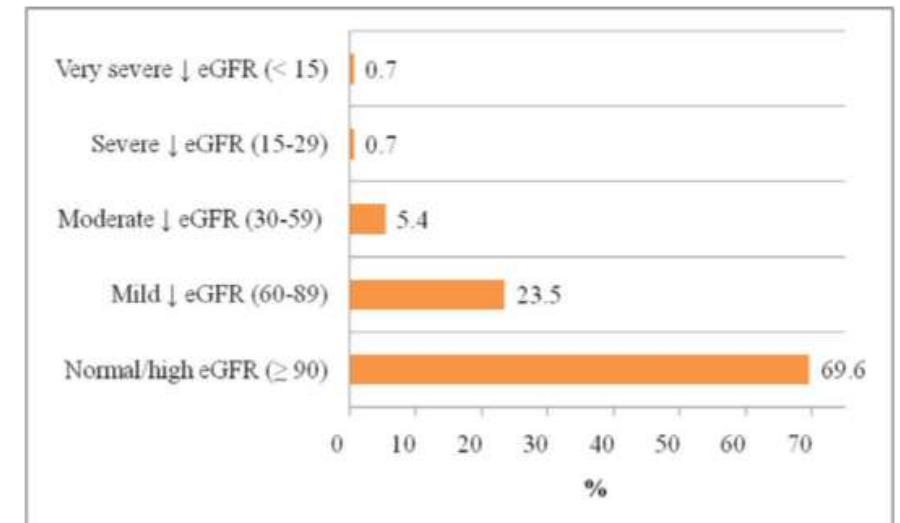
- Started by initiating injury
- Recruit inflammatory cell
- Release growth factor (balance between good and bad)
- Metalloproteinase, PA (cut the collagen)
- RAAS system activation: Ag II pro fibrotic

CKD stages

			Albuminuria categories (mg/g creatinine)		
			A1	A2	A3
			< 30	30–300	> 300
GFR categories (ml/min × 1.73 m ²)	G1	≥ 90			
	G2	60–89			
	G3a	45–59			
	G3b	30–44			
	G4	15–29			
	G5	< 15			
G5 D/T		Kidney failure	Usually defined by KRT		

Epidemiology

- 10% of the adult population around the world have CKD stages 1–5
- Registration and diagnosis (variable between countries)
- found that approximately 31% of the individuals (at high risk) had unrecognised CKD eGFR < 90.
- 7% of the present sample had an eGFR of
- Mortality is higher than non CKD



Risk factors

- Individuals with any of the following attributes are at high risk for CKD and could benefit from screening:
 - 1- age more than 60 years
 - 2- Diabetes mellitus (DM)
 - 3- Hypertension (HTN)
 - 4- Family history of kidney diseases

Leading causes of CKD

- 1- Diabetes
 - 2- Hypertension
 - 3- Obstructive nephropathy (kidney stones, BPH)
 - 4- Kidney diseases (TIN, GN, ADPKD, recurrent UTI and pyelonephritis)
 - 5- Renovascular diseases
 - 6- Some medications - for example, NSAIDs, Heavy metals
 - 7- Fetal developmental problem
 - 8- Infections like Hep C and HIV, Malaria and yellow fever
 - 9- Illegal substance abuse - such as heroin or cocaine.
- Injury - a sharp blow or physical injury to the kidney

Clinical manifestation

- **Urinary:**
 - Urine output: polyuria, oliguria, anuria
 - Frothy, hematuria
- **Uremia:** accumulation of waste product (protein metabolism)
- **Volume:** hypervolemia
- **Acid base, electrolytes:**
 - Sodium
 - Potassium
 - Acidosis
 - Ca
 - Phosphate

- **Hematological**

- Anemia
- Bleeding: platelets dysfunction

- **Infection:**

- **CVS:** CAD, HF, LVH, arrhythmia, pericarditis
- **RS:** pulmonary edema
- **GI:** N/V, ulcer, uremic fetor
- **Neuro:** encephalopathy, peripheral neuropathy, restless leg syndrome

- **Skin**

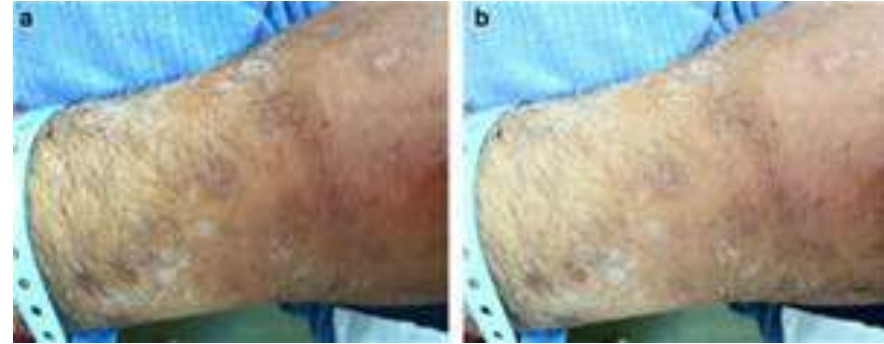
- Earthy color

- Pruritus

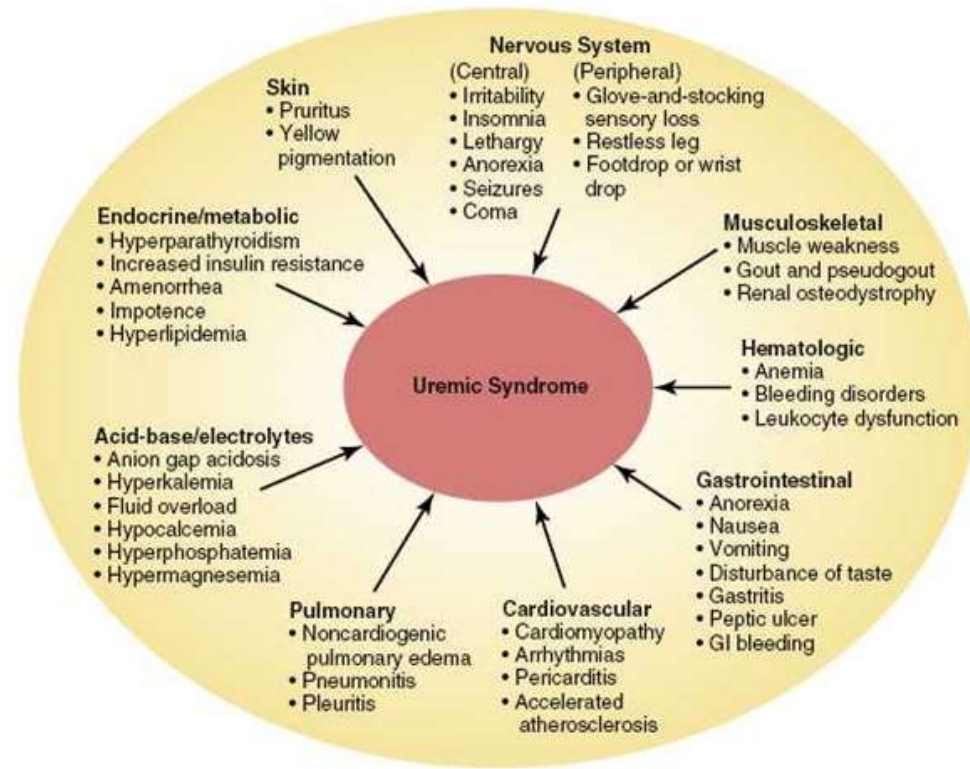
- Petechiae, ecchymosis

- Nail changes

Uremic Forest







Investigation

- Urea/Cr
- Urinalysis and quantification of proteinuria;
- Electrolytes; hyperkalaemia and acidosis, Calcium, phosphate, parathyroid hormone; Albumin
- Full blood count (\pm Fe, ferritin, folate, B12);
- Lipids, glucose \pm HbA1c;
- Renal ultrasound: size, asymmetry, cyst
- Hepatitis and HIV serology
- ECG
- SPEP, serological test

Management

- Acute versus chronic (baseline)
- Reversibility
- Retard progression
- Treat complication
- Prepare for renal replacement therapy

Kidney function

② **Blood filtration**

① Unfiltered blood from body

③ Filtered blood to body

④ Urine with waste and excess fluid to bladder

Prevention of chronic kidney disease

Stop smoking

Reduce salt intake

Eat a heart-healthy diet rich in fruits and vegetables

Exercise regularly and maintain a healthy weight

Control diabetes and monitor HbA_{1c} levels

Control high blood pressure with kidney-protective blood pressure medications

Ureter

Bladder

Kidney

S. Welker

AKI or CKD

	Acute Kidney Injury	Chronic Kidney Disease
History	Short duration History of acute illness	Long duration History of kidney disease and/ or causative co-morbidity
Examination	Acutely ill May have hypotension, fluid overload, metabolic acidosis,	Better toleration of biochemical abnormalities Anaemia, cachexia, grey discolouration of skin
Creatinine	Rapidly increasing values	History indicating derangement Stable serial measurements
Calcium	Usually normal	Low in untreated chronic disease
Haemoglobin	May be normal or low	Due to erythropoietin deficiency- chronic normocytic anaemia
Renal ultrasound	Often normal When the cause is acute obstruction- hydronephrosis	Scars from reflux nephropathy Renovascular disease- asymmetry Large kidneys in • Diabetes, • Amyloidosis, • Chronic Hydropnephrosis • Polycystic Kidney disease

Adapted from Hall (2008)

Treat the cause

- GN
- Pyelonephritis
- AIN
- Obstruction
- Renovascular disease

Progression

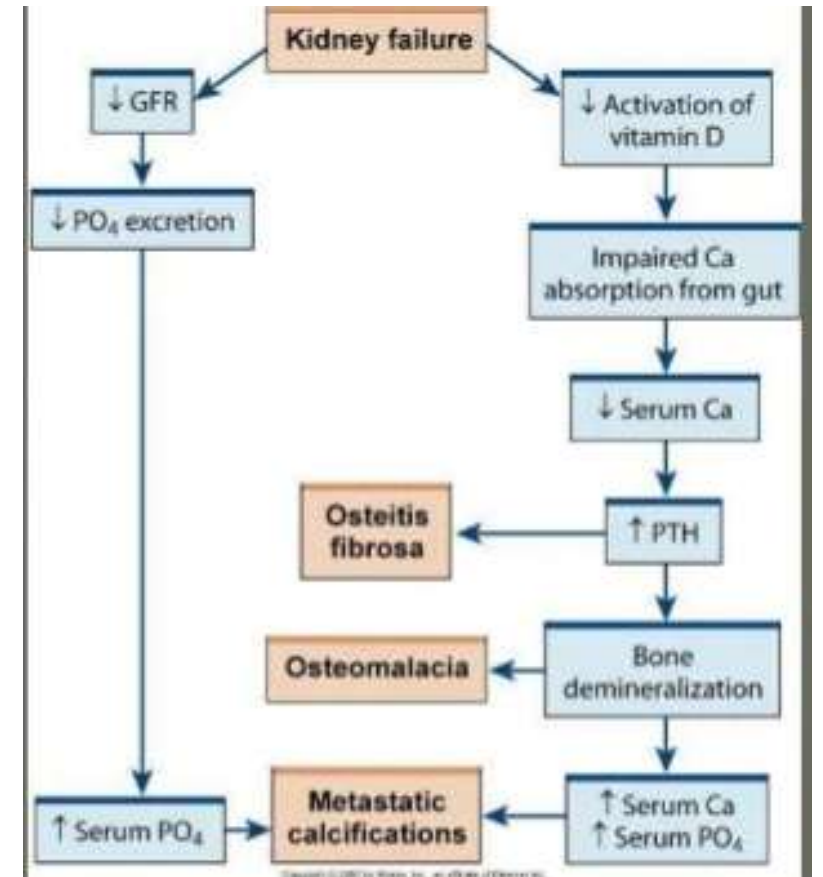
- BP
- Glycemic control
- RAAS blocker
- SGLT2 inhibitors
- Stop smoking
- Diet: protein, salt, phosphate, K
- Exercise
- Obesity
- Acidosis $\text{HCO}_3^- < 22$
- Uric acid ?
- Dyslipidemia

Table 33-4 Drug Dosages in Chronic Kidney Disease

Major Dosage Reduction	Minor or No Reduction	Avoid Usage
Antibiotics		
Aminoglycosides	Erythromycin	
Penicillin	Nafcillin	Nitrofurantoin
Cephalosporins	Clindamycin	Nalidixic acid
Sulfonamides	Chloramphenicol	Tetracycline
Vancomycin	Isoniazid, rifampin	
Quinolones	Amphotericin B	
Fluconazole	Aztreonam, tazobactam	
Acyclovir, ganciclovir	Doxycycline	
Foscarnet		
Imipenem		
Others		
Digoxin	Antihypertensives	Aspirin
Procainamide	Benzodiazepines	Sulfonylureas
H ₂ antagonists	Quinidine	Lithium carbonate
Meperidine	Lidocaine	Acetazolamide
Codeine	Spirolactone	NSAIDs
Propoxyphene	Triamterene	Phosphate-containing bowel-preps

Complication

- Treat anemia target (10-11)
- Secondary hyperparathyroidism
- CV disease



RRT

- Transplant
- Hemodialysis
- Peritoneal dialysis
- Artificial kidney
- Animal kidney ?