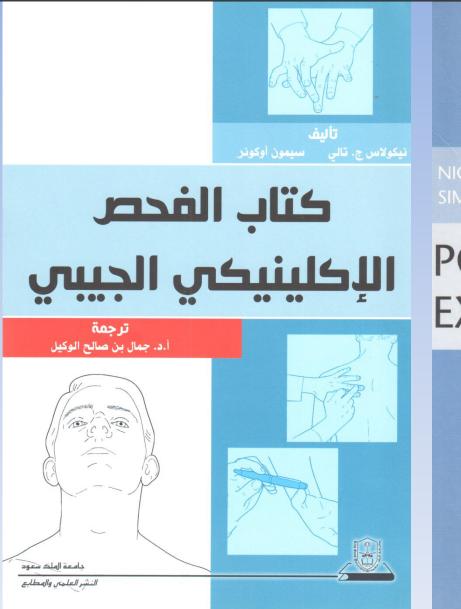


Prof. Jamal Al Wakeel

Professor of Internal Medicine, Nephrology
Notes by Ghaida Alsaeed Consultant

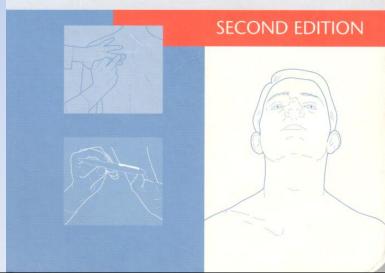
Nephrology Unit, Department of Medicine





NICHOLAS J TALLEY SIMON O'CONNOR

POCKET CLINICAL EXAMINATION



Chronic Kidney Disease

Objective

- Epidemiology of CKD
- Definition of CKD
- Classification
- Symptoms, signs and complications
- Management of CKD

What is the prevalence of chronic kidney disease?

- a. From 1 % to 3 %
- b. From 10% to 16%
- c. From 5 % to 8%
- d. Age dependent

Ans: D

Prevalence of CKD

- CKD in aged ≥20 yrs is >10% -16%
- The prevalence of CKD increases with age:
- 4% at age 29-39 y
- 47% at age >70 y
- The majority of cases are stage 3
- mortality in patients with CKD was 56% greater than that in patients without CKD.
- The 5-year survival rate for a patient undergoing longterm dialysis in the United States is approximately 35%
 - Normally by age, each 1 year increase, GFR decrease 1mL.
 By age of 80, GFR decrease 40 mL

Patients in replacement therapy in SA in 2016

Tx. Pts 2017 11,509

Total HD Pts. 2017 4 | Page : 1012-1020

In 2017, Peritoneal Dialysis a total of 1,389 patients

2016 Total No 26225 pt

Year : 2018 | **Volume** : 29 |

Issue:

2017 Total No 31168 pt

mean total cost of dialysis per patient per year was 46,332 USD (173,784 SR)

Total 1,215,056,700 SR *

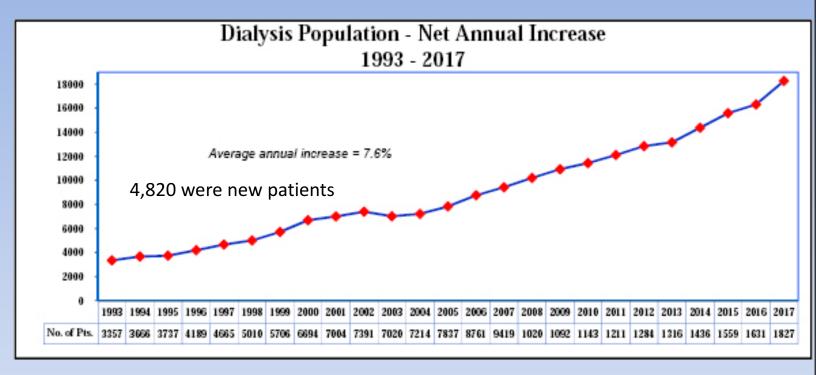
Kalid Al Saran, Alaa Sabry

*audi J Kidney Dis Transpl 2012;23(1):78-82 © 2012 Saudi Center for Organ Transplantation

4.7% increase in new hemodialysis patients. in 2016

Costy disease.

Best treatment of CKD is prevention



• Dialysis in the Kingdom of Saudi Arabia. Saudi J Kidney Dis Transpl [serial online] 2018 [cited 2020 Mar 17];29:1012-20. Available from: http://www.sjkdt.org/text.asp?2018/29/4/1012/239666

Risk Factors CKD

- 45% of dialysis patients have DM
- 35% of dialysis patients have HTN

Which of following are risk factors for CKD (more than one answer)??

- TB infection
- History of colon cancer
- Smoking
- History of SLE
- Obesity

Risk Factors CKD

Diabetes Mellitus 30%

30% of DM patients reach ESRD

Hypertension 25%

25% of HTN patients reach ESRD

Old age 50%

Low GFR E.g. small kidney
AKI

More susceptible to CKD

Obesity

Compared to HTN and DM they play the same role of leading to stroke, ESRD

Cardiovascular disease

E.g. Congestive heart failure, ischemic heart disease. Cardiorenal syndrome: meaning cardio leads to renal

failure

NSAID

Family History

Familial diseases like polycystic kidney disease, alport syndrome are more susceptible

Smoking

Smokers have HTN and vasoconstriction

Table 4.8. Causes of end-stage renal disease in HD patients 2016						
Total	16,315	100%				
Others	673	4%				
Pregnancy Related	73	0.4%				
Primary Glumerular Disease	128	1%				
Vasculitis	215	1%				
Congenital Malformation	271	1.6%				
Hereditary Renal Disease	279	2%				
Obstructive Uropathy	339	2%				
Primary Tubulo Instertitial Disease	491	3%				
Unknown Etiology	1100	7%				
Hypertensive Nephropathy	6211	38%				
Diabetic Nephropathy	6535	40%				

Cause of Renal Failure

By preventing diabetic and hypertensive nephropathy number of CKD patients decrease by 80%.

chronic Kidney disease can be diagnosed??

- a. If the patient have ultrasound shows scaring in the kidney
- If patient have only proteinuria but normal serum Cr
- If the patient have high serum creatinine.
- If the patient have proteinuria and hematuria with low eGFR

Answer is discussed in the next slides

Chronic kidney disease is a new term we used to call it chronic renal failure, renal impairment, and impaired renal function.



Kidney International Supplements 2017 7. 1-59DOI: (10.1016/i.kisu.2017.04.001



International Supplements 2017 7, 1-59DOI: (10.1016/j.kisu.2017.04.001

Any single change (either structural or functional) for 3 months or more is a chronic kidney disease

Structural (kidney shape):

- -Ultrasound: scarring
- -Biopsy: e.g. glomerulonephritis

Kidney damage

Functional:

- -Proteinuria
- -Hematuria
- -Electrolyte imbalance
- -High serum creatinine
- -Acidosis

GFR normal range depends on the age but it range from 110-140 Structural abnormalities

Structural abnormalities detected by imaging

Abnormalities detected by histology
History of kidney transplantation

Functional abnormalities

Albuminuria

Electrolyte and other abnormalities

Urine sediment abnormalities tubular disorders

GFR < 60 ml/min/1.73 m²

for ≥3 months

Chronic kidney disease definition: A disease that damage the kidney and decrease the function over 3 months presented by GFR < 60

Staging of chronic kidney disease depend on ?

- a. Measurement of serum creatinine
- b. Measurement creatinine clearance
- c. Estimated GFR
- d. Urine albumin creatinine ratio
- e. Causes of chronic kidney disease

Ans: C.

- +E? because doctor said each disease has a different outcome for example DM is faster than HTN, so you have to mention the cause with the stage.
- DM takes 5 years to reach ESRD.
- HTN takes 5-10 years to reach ESRD.
- Glomerulonephritis takes 20 years

KDIGO recommendation -classified based on:(CGA)

- Cause
- –Stage of CKD (GFR category)
- Albuminuria category

Staging depends on 2 things:

- GFR
- Protienuria

KDIGO recommendation -classified based on:(CGA)

- Cause
- -Stage of CKD (GFR category)
- Albuminuria category

CKD Classification Estimation and Measurement of GFR

Estimate and/or Measure GFR

Filtration Markers (Endogenous)

Currently we do this.
It's a calculated equation,
it depends on patient data

eGFR equations

eCKD- EPI cr

eCKD-EPI-cyc

Creatinine Clearance (CrCl)

24 h collection of urine. Normal range is 110-140 Cystatin C
Polypeptide

chain of 120 amino acids

If high > CKD, If low > normal

Unlike creatinine, higher level of cystatin c polypeptide is <u>not</u> correlated with worse CKD. It's just a marker of CKD.



Chronic Kidney Disease Epidemiology Collaboration (CKD-EPI)

Modification of Diet in Renal Disease (MDRD)
Study equation

Calculators

There are many formulas to calculate eGFR:

The first 2 formulas are the most commonly used

CKD-EPI Creatinine Equation

Preferred method

MDRD Study Equation

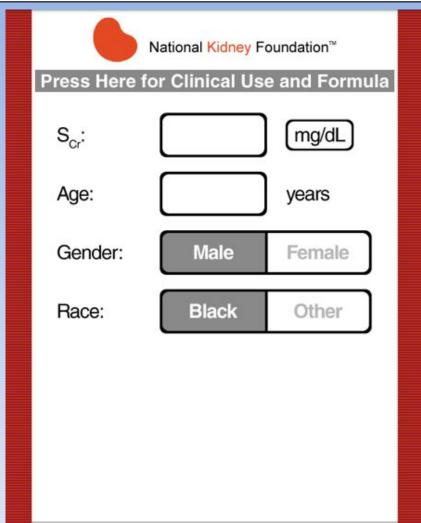
CKD-EPI Cystatin and Creatinine 2012 Equation

Cockcroft-Gault Formula

Revised Bedside Schwartz Formula

For ages 1 - 17

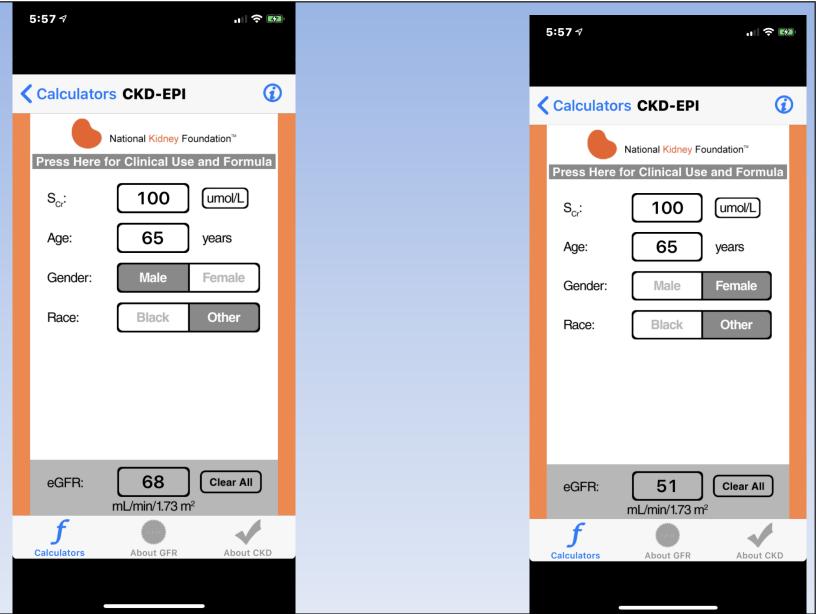
CKD — EPI Calculation



mL/min/1.73 m²

eGFR:

Clear All



Staging

The stages of CKD are classified as follows

- Stage 1: Kidney damage with normal or increased GFR (>90 mL/min/1.73 m²) Normal or high High risk patients: they have normal GFR with proteinuria or hematuria or acidosis or hyperkalemia or structural changes (any structural or functional abnormality). When treated early, CKD is prevented.
- Stage 2: Mild reduction in GFR (60-89 mL/min/1.73 m²) Mildly decreased*
 Slow progression, correct the risk factors, they will not go to CKD
- Stage 3a: Moderate reduction in GFR (45-59 mL/min/1.73 m²) 3a: Slow progression
- Stage 3b: Moderate reduction in GFR (30-44 mL/min/1.73 m²) 3b: Rapid deterioration of kidney function
- Stage 4: Severe reduction in GFR (15-29 mL/min/1.73 m²)
- -Symptoms start to appear when GFR < 45 with nephrotic syndrome.
- -without nephrotic syndrome symptoms appears
- Stage 5: Kidney failure (GFR < 15 mL/min/1.73 m² or dialysis) when GFR < 30 Stage 5 patient need to start dialysis (due to symptoms of uremia)

Stages of chronic kidney disease 1 3b 4 5 Normal or Severely Kidney higher decreased kidnev function function GFR > 90 GFR 29-GFR 44-30 15

The majority of cases are stage 3

KDIGO recommendation -classified based on:(CGA)

- Cause
- Stage of CKD (GFR category)
- Albuminuria category

Albuminuria categories as follows:

*note that where albuminuria measurement is not available, urine reagent strip results

Differentiate between proteinuria and albuminuria can be substituted

When patient has 30 albuminuria, his proteinuria would be 50 (albumin + other proteins)

Here we're talking about albuminuria

Albuminuria categories in CKD

Category	AER (mg/24 hours)	ACR (mg/mmol)	ACR (mg/g)	TERMS
		approximate	equivalent	
A1	< 30 No proteinuria to mild proteinuria	<3	<30	Normal to mildly increased
A2	30-300 They used to call it micro albuminuria	3-30	30-300	Moderately increased*
А3	>300 Here proteinuria would be because albuminuria is 3		>300	Severely increased**

^{*}Relative to young adult level

[ACR .2220/g; >220 mg/mmol])1

^{**} Including nephrotic syndrome (albumin excretion usually > 2200 mg/24 hours

Prognosis of CKD by GFR and albuminuria categories: KDIGO 2012		Normal to mildly increased <30 mg/g <3 mg/mmol	Moderately increased 30–300 mg/g 3–30 mg/mmol	Severely increased >300 mg/g >30 mg/mmol	The graph shows you the outcome of CKD. For example when patient is stage 1 (GFR > 90) with A1 (albuminuria < 30) he has a very low risk.				
categories (mL/min/1.73 m²) description and range	GI	Normal or high	≥90				Another example when patient is stage 1 (GFR > 90) with A2 (albuminuria 300) he has increased risk.		
	G2	Mildly decreased	60–89				He will take 12-15 years to develop CKD		
	G3a	Mildly to moderately decreased	45–59						
	G3b	Moderately to severely decreased	30–44						
	G4	Severely decreased	15–29						
GFR	G5	Kidney failure	<15						
otes: Green indicates low risk (if no other markers of kidney disease, no CKD); yellow indicates moderately increased risk; orange indicates high risk; red indicates gh risk. Reprinted with permission from Macmillan Publishers Ltd: Kidney Int Suppl. 2013;3:1–150. Kidney Disease Improving Global Outcomes (KDIGO). 2012 cl									

Another example when patient is stage 1 (GFR > 90) with A2 (albuminuria 300) he has increased risk. He will take 12-15 years to

Persistent albuminuria categories description and range

ange indicates high risk; red indicates very high risk. Reprinted with permission from Macmillan Publishers Ltd: Kidney Int Suppl. 2013;3:1-150. Kidney Disease Improving Global Outcomes (KDIGO). 2012 clinical

practice guidelines for the evaluation and management of chronic kidney disease. Copyright © 2013.15 Abbreviation: CKD, chronic kidney disease.

History

You have to take a full hx

- History of major complain
- History of present illness
 - Upper urinary tract
 - Lower urinary tract
 - History of uremia
 - Systemic review (arthritis ,ch infection ,hepatitis)
 - History of hypertension(cause, duration, control, medication, complication)
 - Diabetes Mellitus(type, duration, control, medication, complication)

History

- Systemic review
- Past Medical illness+ hospitalization
- Surgical
- Medication
- Family
- Social
- Allergy+ medication side effect

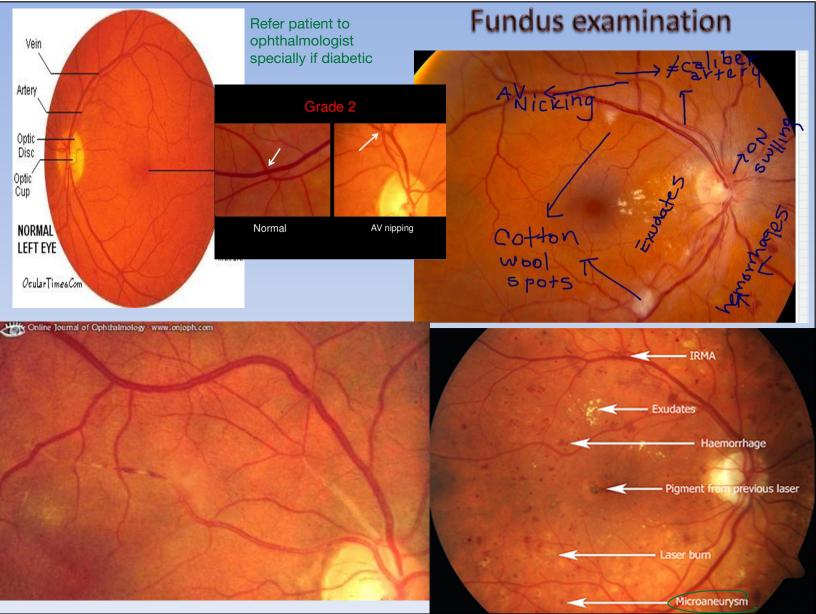
Examination

Full examination is essential because it affects the body from head to toe

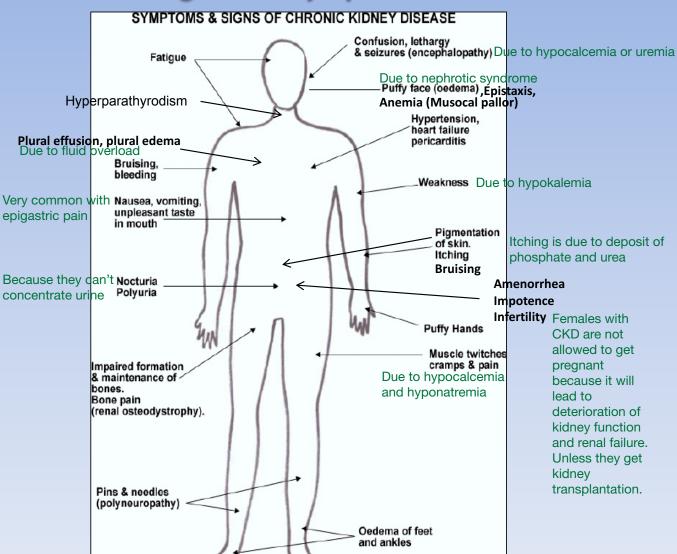
- General look
- Vital sign
- General examination
- Cardiovascular JVP
- Respiratory
- Abdominal
- CNS

Edema always found on the medial malleolus





Signs and Symptoms



Which of the following is essential investigation for CKD?

- Urine electrolyte
- pT, PTT, INR
- Calcitonin & parathyroid hormones
- Kidney Biopsy
- Na,K,Ca, Phosphorus,urea and Cr

Investigations of CKD

- Investigations of diagnosis of CKD
- Investigations to diagnose the cause of CKD
- Investigation to diagnosis the complications of CKD

Investigations of CKD

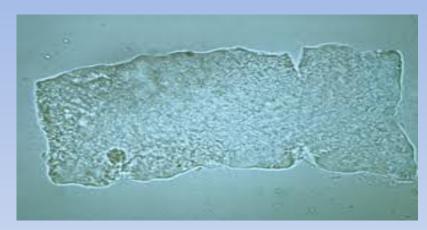
Basic Laboratory to the diagnosis of CKD can include the following:

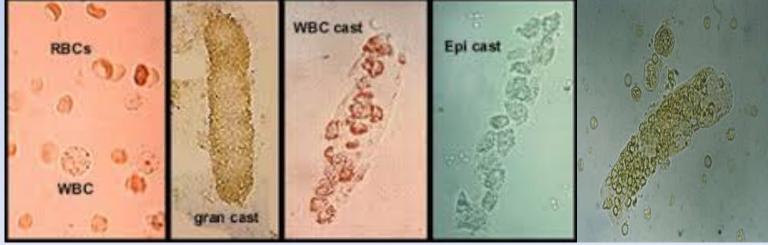
- Complete blood count (CBC) To check anemia
- Biochemistries (Na, K, Urea, Cr, HCO₃, Ca, PO3, Uric, Albumin, Alk ph)
 Hypoalbuminemia > nephrotic syndrome

 Serum creatinine is essential to measure GFR, bicarbonate is imp to check acidosis
- Urinalysis Imp because sometimes it's enough to make the diagnosis
- 24 hour urine collection Creatinine clearance and proteinuria To classify the patient to A1, A2, or A3
- Urine albumin:creatinine ratio
- Glucose
- Lipid profile: increased risk of cardiovascular disease



الكاست عباره عن بروتين





Investigations of CKD

Laboratory studies used in the diagnosis the cause of CKD

HbA1c

ANA To rule out SLE

HBsAg

HCV. Causes silent CKD

 C_{3} , C_{4} Some glomerulonephritis patients have low c3, c4

Renal Biopsy

CT, MRI, and Radionuclide Scans

• Renal Ultrasonography: initial imaging US is essential

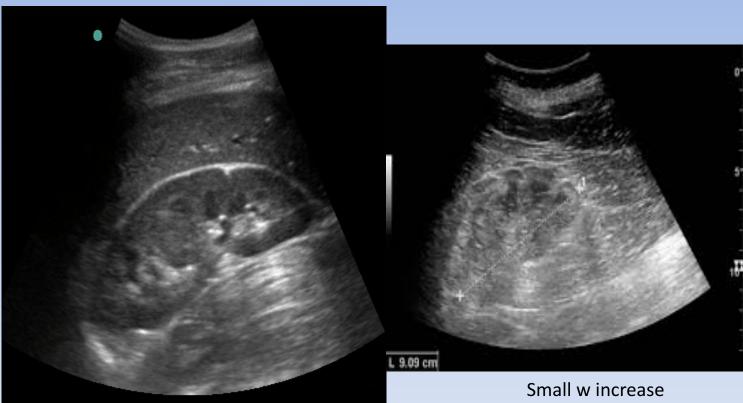
- CT, MRI: Intravenous (IV) should be avoided

 Because it causes more renal damage
- CT & MRI only in case of obstruction or renal stone
- Radionuclide Scans: assist GFR & obstruction
 Tc^{99m}MAG3 and .Tc^{99m}DTPA.Tc^{99m}DMSA

Choose the type of radionuclide scan depending on the disease:

- DMSA > kidney scarring
- MAG3 + DTPA > measure renal function, GFR

Investigation



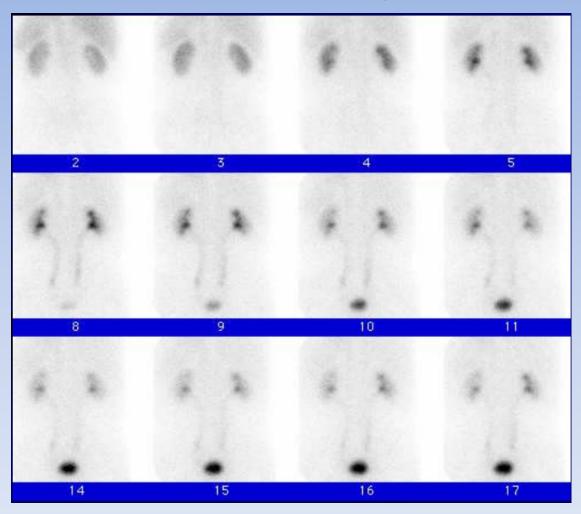
Corticomedullary preserved

- •Normal size kidneys 12 cm
- Normal Echogenicity

Small w increase echogenicity
Compared to liver echogenicity

Shrunken + corticomedullary lost

Nuclear scan safe in kidney disease



Investigation to the diagnosis the complications of CKD

1- CBC: to check anemia



Bone metabolic disease or mineral metabolic disease



Most renal patients have low vit D because it's lost with the protienuria. Vit D can reduce blood pressure, improve cardiac disease and other benefitial metabolic effect.



Because many patients have valvular calcification, vessels calcification and HTN which leads to LVH



Diagnosis of CKD

- Different
 - ✓ Acute Kidney Injury form
 - **✓** CKD

Check patient file if he has any renal abnormalities before 3 months or more.

Differentiating between acute and chronic kidney disease

- **1- Proper hx:** including family hx (e.g. polycystic kidney disease)
- 2- PE: S4 heart sound on auscultation, eyes (hypertensive or diabetic retinopathy), renal bruit > chronic
- **3- Biochemistry:** hypocalcemia + hyperphosphatemia > chronic **CBC:** low hemoglobin > chronic
 - **Urinalysis:** RBC cast, WBC cast > Acute, broad widely cast > chronic
- **4- Ultrasound:** if shrunken kidney > chronic. If enlarged kidney > acute
- 5- Check for any exacerbating factors, correct any reversible factors like drugs.

Treatment of chronic kidney disease??

- a. The treatment usually able to cure chronic kidney disease
- b. Its slow progressive of chronic kidney disease
- c. It include treatment of hypertension and reduce proteinuria
- d. Alkalosis should be treated in chronic kidney disease

We're looking for a world without kidney disease in the next 20 year

Diet:

High protein diet is nephrotoxic.

PO3 is used to preserve food but it's toxic for the kidney.

Patients should eat fresh food with low sodium and high potassium to protect the kidney and prolong age

> Prevention &Slow progress of CKD



Avoid Nephrotoxic Drug **Glycemic Control**

BP control

Proteinuria & RAAS

Give ACE + ARBs to treat proteinuria

Diet

Protein, PO3 & salt

Correct Acidosis

Hyperlipidemia

Plus contrast and many herbal medicine

Anyone above 50 y should take statins even if lipid profile is normal

Target Blood Pressure in CKD

≤130/80

Diabetics and Non-diabetics				
Albuminuria		Drug of choice		
A1 <30 mg/d		CBC, Diuretic, RAAS		
A2 >30 mg/d		ARB or ACE-I		

If +ve proteinuria, you must give ACE inhibitors + ARBs

If -ve proteinuria, you can give any hypertensive agent e.g. diuretics, CCB.

Better to use small doses + combination so you get the benefit of each drug with minimal side effects.

No B-blocker

Reduce proteinuria

- Low salt diet Because salt increases proteinuria
- Inhibit renin-angiotensin system (RAS) by Angiotensin-Converting-Enzyme Inhibitor (ACE Inhibition) or angiotensin-II-receptor blocker (ARB)

Check serum creatinine and K after 1 week, if not increased continue the medications

Renal Diet

- Low salt diet 2.4-3 g/day Daily requirement in healthy people is 5-8. Zero salt diet is wrong it might increase the complications
- Low potassium diet For patients on ACEI + ARBs with increased K.
 - o (2g-2.5 g), half of normal intake
- Low protein diet 0.6 to 0.8 g/kg per day Our need is 0.6
- Low phosphorous diet 800 1000 mg/day High phosphorus is dangerous. Found in red meat and canned food.
- Water intake daily water intake 1.5-2 L
 Avoid excessive water it can lead to hyponatremia
- Restricted Magnesium

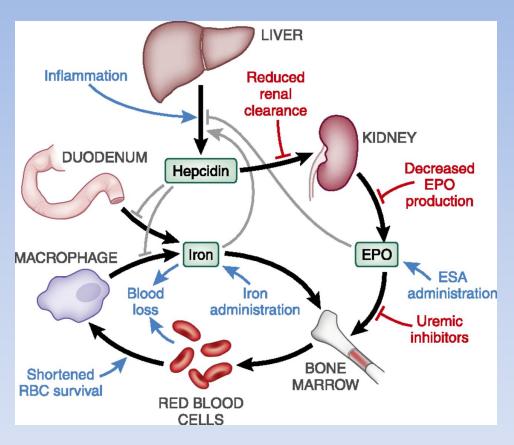
Treatment of Complications

- Volume overload: Which leads to pulmonary edema and HTN
 - o restrict SALT
 - o loop diuretics
- Metabolic acidosis:
 - oral alkali supplementation

We give Sodium bicarbonate (NaHCO₃). Unlike Sodium chloride, it doesn't increase the pressure.

Anemia in CKD

Anemia is common in patients with GFR < 45.



Anemia happens due to:

- Decreased erythropoietin (EPO).
- Hepcidin (inflammatory marker) released by liver, it decreases iron absorption.
- Shortened RBCs life span in CKD from 120 to 35-40 Causes Decrease EPO Increase Hepcidin- iron Bone marrow suppression Shortened RBC survival



In chronic diseases, high **hepcidin** production inhibits iron release from macrophages and intestinal absorption of iron. This consequently induces an **anemic** condition.

Treatment of anemia

- Anemia: hemoglobin level \$\square\$10 g/dL check iron -iron Tablet or IV erythropoiesis-stimulating agents (ESAs)
 - ✓ epoetin alfa
 - ✓ darbepoetin alfa
 - ✓ Methoxy polyethylene glycol-epoetin beta(Mircera)

Normal ferritin levels range from 100-200. In CKD, we want ferritin levels to reach 300-400. We give iron tablets or IV iron, if not corrected we give ESAs الكعمل، واحد يعطى يوميا، واحد اسبوعيا، واحد شهريا. يعطى حسب هيموقلوبين المريض ESAs النامة increases the risk of stroke and malignancy, should be avoided in high risk patients

Hyperkalemia

Very important to measure in patients on ACEI + ARBs

- Cause :
- Reduce GFR
- ARBs (angiotensin II receptor blockers). ...
- ACE (angiotensin converting enzyme) inhibitors....
- Spironolactone
- High K diet مثل التمر، التين، المشمش، المنقا ،الموز والطماطم

Asymptomatic & weakness & Arrythmia

Hyperkalemia is a medical emergency

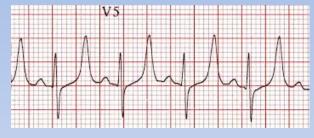














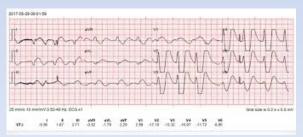


Table 1. Emergency management of acute hyperkalemia

Medication	Response Type	Onset of action	Duration of action	Mechanism Exp	ected decrease in potassium level
Calcium gluconate	rapid	1-2 min	30-60 min	Protect cardiomyocytes	
Glucose + insulin	intermediate	10-20 min	2-6 hours	Shift potassium intracellula	orly 0.5-1.5 mEq/L (dose dependent)
Beta-agonist	intermediate	3-5 min	1-4 hours	Shift potassium intracellula	ırly

2-6 hours

4-6 hours

2-6 hours

Shift potassium intracellularly

Elimination of potassium from the body

Elimination of potassium from the body

Elimination of potassium from the body

1mmol/L in the first 60 min and total of

2mmol/L by 180 min

It doesn't manage hyperkalemia, it only protects the heart against hyperkalemia. In ESRD (CKD) those medications don't work. Manage them with dialysis

30-60 min

2-6 hours

5-30 min

immediate

Sodium bicarbonate

Exchange resin

Furosemide

Hemodialysis

intermediate

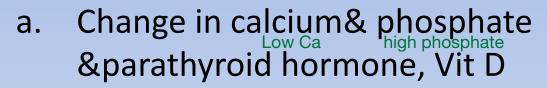
delayed

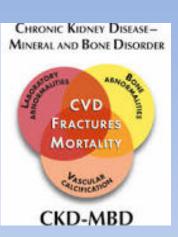
delayed

delayed

CKD MBD indicate

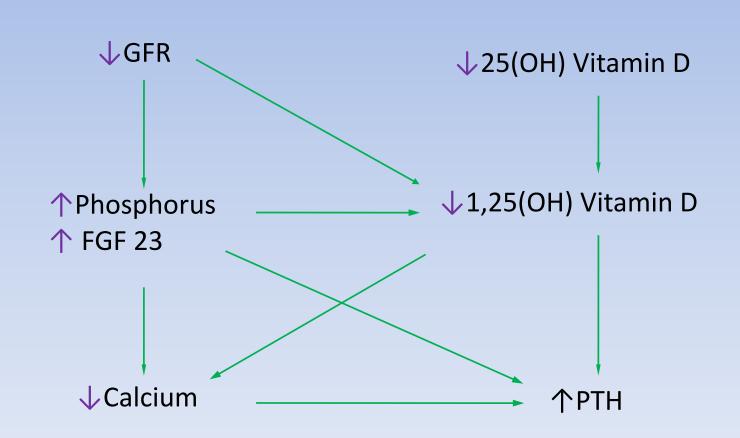
Mineral & Bone Disorder





- b. change in the bone metabolize and mineral
- Indicate change in the extra skeletal calcification and vascular, valvular calcification
- Ca + phosphate normally deposit in bone but in CKD MBD they deposit in other tissues like blood vessels wall (vascular calcification) and valve (valvular calcification).
- MBD affect 3 things: bone (osteodystrophy), blood vessels (calcification), and minerals imbalance (low ca, high phosphorus, and high parathyroid).

Mineral abnormalities of Chronic Kidney Disease (CKD)



Renal osteodystrophy

- high bone turnover disease related to secondary hyperparathyroidism (referred to as osteitis fibrosa cystica), Makes the bone more prone to fracture with minor trauma.
- low turnover disease (referred to as adynamic bone disease),
 - Patients with low vitamin D and hyperparathyroid, when givin high vit D supplements it will suppress parathyroid and lead to **adynamic bone disease**.
- osteomalacia (low turnover disease accompanied by undermineralized bone tissue) osteomalacia occurs in children with vitamin D deficiency even without CKD.
- Osteoporosis
- mixed disease where features of both high and low bone turnover disease are present

High bone turnover disease



osteitis fibrosa cystica),



AP radiograph of the hand in a 66-year-old woman with primary hyperparathyroidism owing to parathyroid adenoma shows subperiosteal bone resorption (arrows) along the radial aspect of 2nd, 3rd, and 4th middle phalanges.









a. extra skeletal calcification and vascular, valvular calcification

CKD-MBD

Hyperphosphatemia

- √ dietary phosphate restriction
- ✓ dietary phosphate binders (Ca Carbonate, Ca Acetate, Sovlomier, lanthium)

Hypocalcemia

✓ calcium supplements with or without calcitriol

Hyperparathyroidism

- ✓ Calcitriol
- √ vitamin D analogs

Treatment of Complications

• Uremic manifestations:

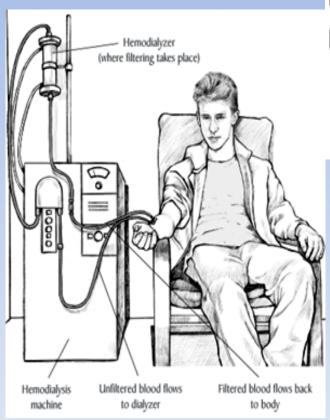
- Long-term renal replacement therapy
 - ✓ hemodialysis
 - ✓ peritoneal dialysis
 - ✓ renal transplantation

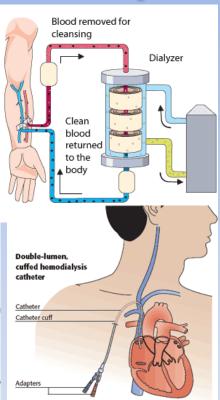
Renal transplantation is better than hemodialysis

Indications for renal replacement therapy

- Severe metabolic acidosis
- Hyperkalemia After management failure.
- Intractable volume overload After management failure.
- Pericarditis
- Uremic Symptoms E.g. Flapping tremors
 - ✓ Encephalopathy✓ Failure to thrive and malnutrition
 - ✓ Peripheral neuropathy
 - ✓ Intractable gastrointestinal symptoms
- In asymptomatic patients
 - ✓ GFR of 5-8 mL/min/1.73 m²

Hemodialysis



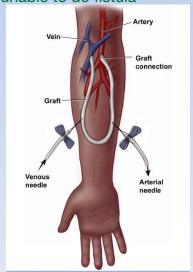






FISTULAS FROM DIALYZER ACCESS FISTULA TO DIALYZER ARTERY

This's graft used when unable to do fistula



Hemodialysis Access:

- 1- Fistula: Needs to be done 6 months before dialysis
- 2- Graft: Needs to be done 2 months before dialysis
- 3- **Catheter**: Use in <u>urgent</u> cases for dialysis to avoid clotting. It's placed in large blood vessels e.g. internal jugular vein, subclavian

Complication of hemodialysis:

- Infection
- Septicemia
- · Clotting. need to do another fistula

Complication of peritoneal dialysis:

- Peritonitis
- · Hemodialysis is 3 times per week. Peritoneal dialysis is 4 times daily
- البيرتونيال ديالسز بيشنتس عندهم حرية الاكل والسفر عكس الهيموديالسز

Renal transplantation:

New kidney is transplanted in the pelvic area. It has the best outcome.

Complications:

- rejection
- More susceptible for infection

Peritoneal Dialysis

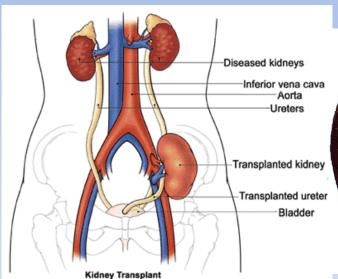






Renal Transplantation

Living related donors (LRD)
Living unrelated donors (LURD)
deceased organ donors







Renal Transplant Medication

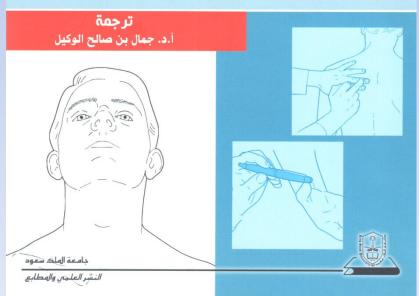
Renal Transplant Rejection& infection

✓ Better qulity of live ,less mortality ,can be before HD



تاليف نيكولاس ج. تالي سيمون أوكونر

كتاب الفحص الإكلينيكي الجيبي





NICHOLAS J TALLEY SIMON O'CONNOR

POCKET CLINICAL EXAMINATION

