

Nephrology Cases and Everything in Between

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Disclosures

- Consulting/Advisory Board for Bayer and Boehringer Ingelheim pharmaceuticals



Objectives

- Differentiate between pre-, intrinsic, and post-renal acute kidney injury (AKI)
- Recognize the differences of presentation between interstitial nephritis (AIN), glomerulonephritis (GN), and nephrotic syndrome
- Interpret the different nephrology lab results that help to guide workup of AKI, ADPKD, ATN, AIN, and GN and electrolyte abnormalities
- Gain confidence in evaluation of altered kidney function



Functions of the kidney

Filtration

Renin-angiotensin
system

Erythropoietin
production

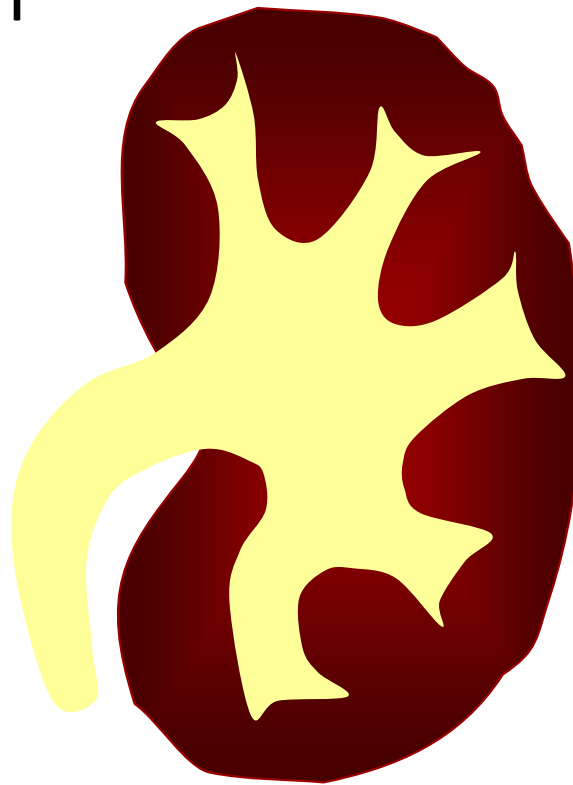
1,25OH-Vit D
production

Free water
homeostasis

Acid-base
balance

Excretion of
uremic toxics

Solute balance



Steps to success in nephrology

1. What is the kidney function? Acute vs chronic
2. What do you see in the urine analysis?
3. How much proteinuria?
4. What do you see on kidney ultrasound?



Urine = success in nephrology

1. IF kidney, look at the urine
2. IF kidney, look at the urine
3. IF kidney, look at the urine



Evaluating kidney function

BUN

- Blood urea nitrogen
- Waste product of protein catabolism
- Normal units 6-20mg/dL
- High BUN found in:
 - high protein diet
 - CKD
- Hypovolemia:
 - CHF, GI hemorrhage, hemorrhage
- Fever
 - Inc catabolism

SCr

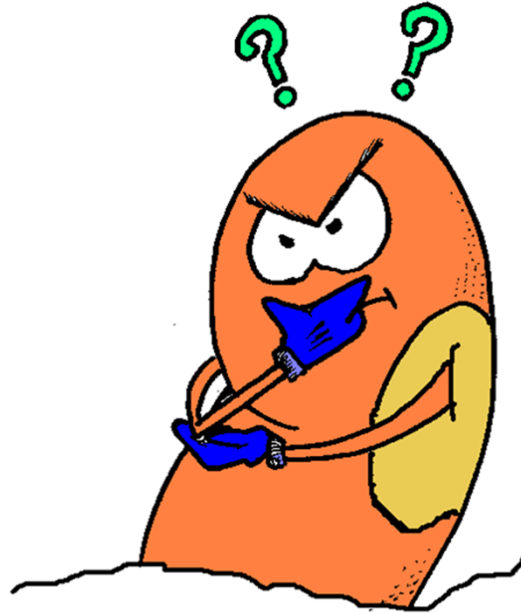
- Serum Creatinine
- Breakdown of creatine phosphate in muscle
- Normal units 0.5-1.2mg/dL
- Dependent on muscle mass
higher muscle mass = higher SCr
- Higher levels in males
- **Can be late marker of kidney function**
- Logarithmic scale



What is the eGFR?

35 y/o male
eGFR 62
Stage 2

45 y/o female
eGFR 43
Stage 3b



70 y/o male
eGFR 50
Stage 3a

80 y/o female
eGFR 35
Stage 3b

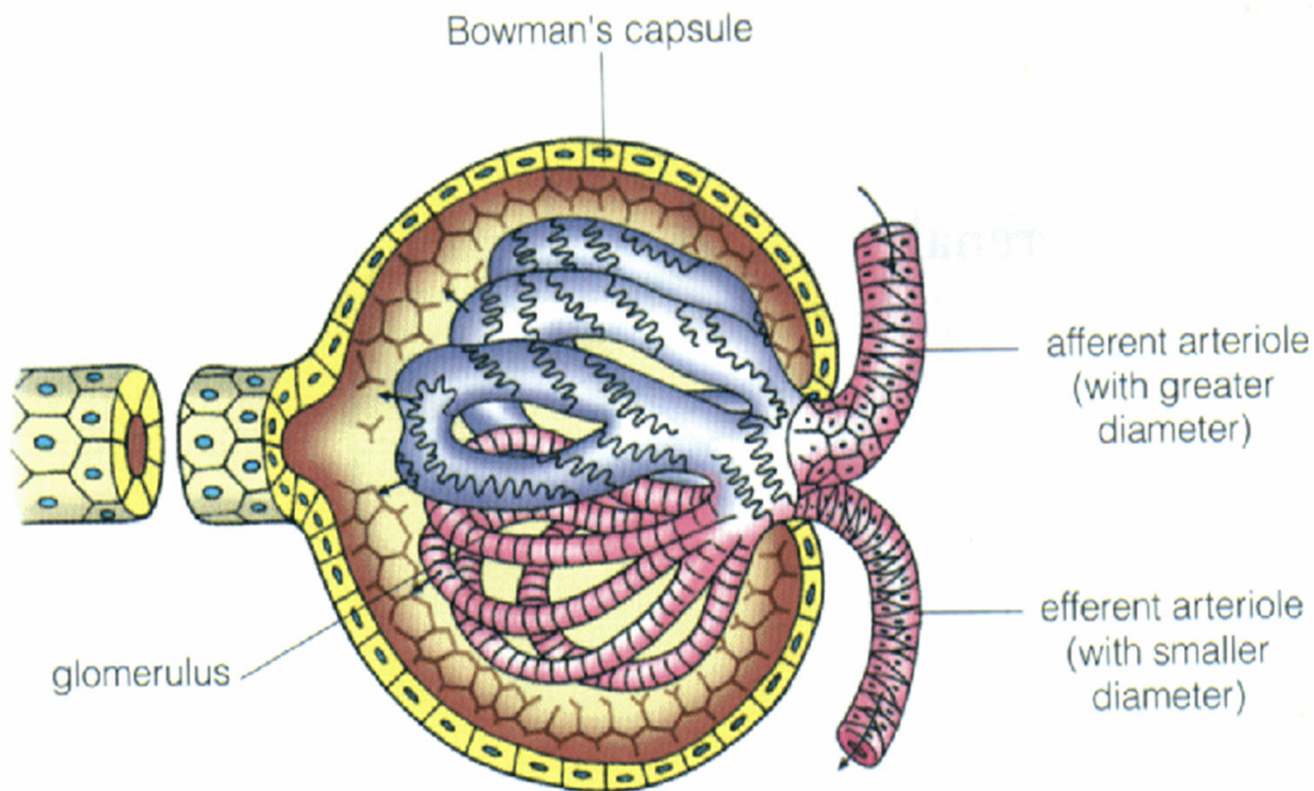
Hold SCr stable at 1.5



Evaluating kidney function

GFR

- GFR measures amount of plasma filtered across glomerular capillaries (nl >90ml/min)



Evaluating kidney function **ESTIMATES** of GFR

MDRD vs CrCl vs CKD-EPI vs CG

eGFR = % of remaining kidney function

- Cr Cl = 24 hour urine collection = clearance of kidneys = estimate of GFR CrCl = often used in FDA inserts, interchangeable with GFR,
- Cockcroft-Gault (CG) = FDA/pharmacy uses CG for med renal dosing = estimate of CrCl
- MDRD = eGFR = based upon age, sex and SCR. NOT accurate when eGFR >60ml/min
- CKD-EPI = **latest eGFR calculation**. More accurate when eGFR >60ml/min
- **GFR requires stable function, not useful in AKI**
- **An increase in SCr from 0.6 to 1.2 indicates a 50% loss of kidney function**





George

80 y/o male

PMH: HDL, HTN, BPH, CAD, former smoker

Meds: atorvastatin, chlorthalidone, amlodipine, tamsulosin, ASA

Retired military who still works full time @ golf shop

Presents to ED with gross hematuria x 1 week. Denies dysuria, fever/chills, penile discharge, LE edema, or changes in urinary habits

Which SCr (mg/dl) change represents most significant deterioration of George's kidney function?

- A. 0.6 → 1.4?
- B. 1.0 → 1.5?
- C. 1.5 → 2.0?
- D. 2.0 → 2.5?





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Which SCr (mg/dl) and eGFR (ml/min) change represent most significant deterioration of George's kidney function?

A. 0.6 → 1.4?

GFR 95 → 47

B. 1.0 → 1.6?

GFR 71 → 40

C. 1.7 → 2.8?

GFR 37 → 20

D. 3.0 → 6.0?

GFR 19 → 8





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IF his baseline renal function was SCr of 0.6 and eGFR of 95 and today the SCr and GFR are 1.4 and 47, what is the likely cause of kidney dysfunction?

- A. Acute kidney injury (AKI)
- B. AKI on chronic kidney disease (CKD)
- C. CKD



Criteria for CKD

1. Decreased GFR \leq 60ml/min for **> 3 months**
OR
2. Markers of Kidney Damage for **> 3 months**
 - Albuminuria \geq 30mg/g
 - Urinary sediment abnormalities
 - Ex: microhematuria
 - Electrolyte and other abnormalities due to tubular disorders
 - Structural abnormalities detected by imaging
 - Ex: ADPKD (polycystic kidney disease)
 - History of kidney transplantation





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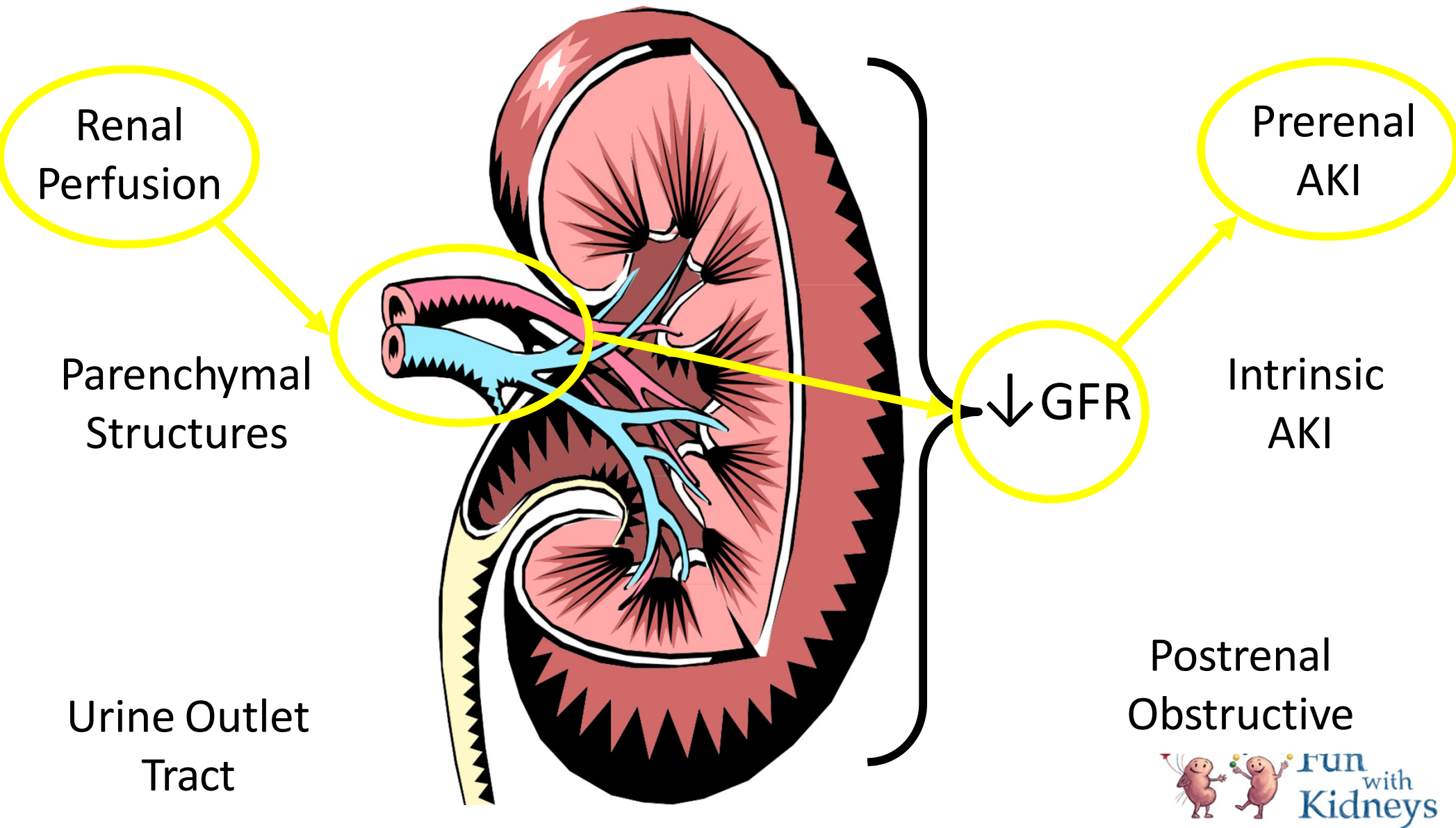


Syndromes of Acute Kidney Injury

Factors that Affect:

Induce:

Syndrome:



Renal
Perfusion

Parenchymal
Structures

Urine Outlet
Tract

Prerenal
AKI

Intrinsic
AKI

Postrenal
Obstructive

↓ GFR

Syndromes of Acute Kidney Injury

Factors that Affect: Induce: Syndrome:

Renal
Perfusion

Prerenal
AKI

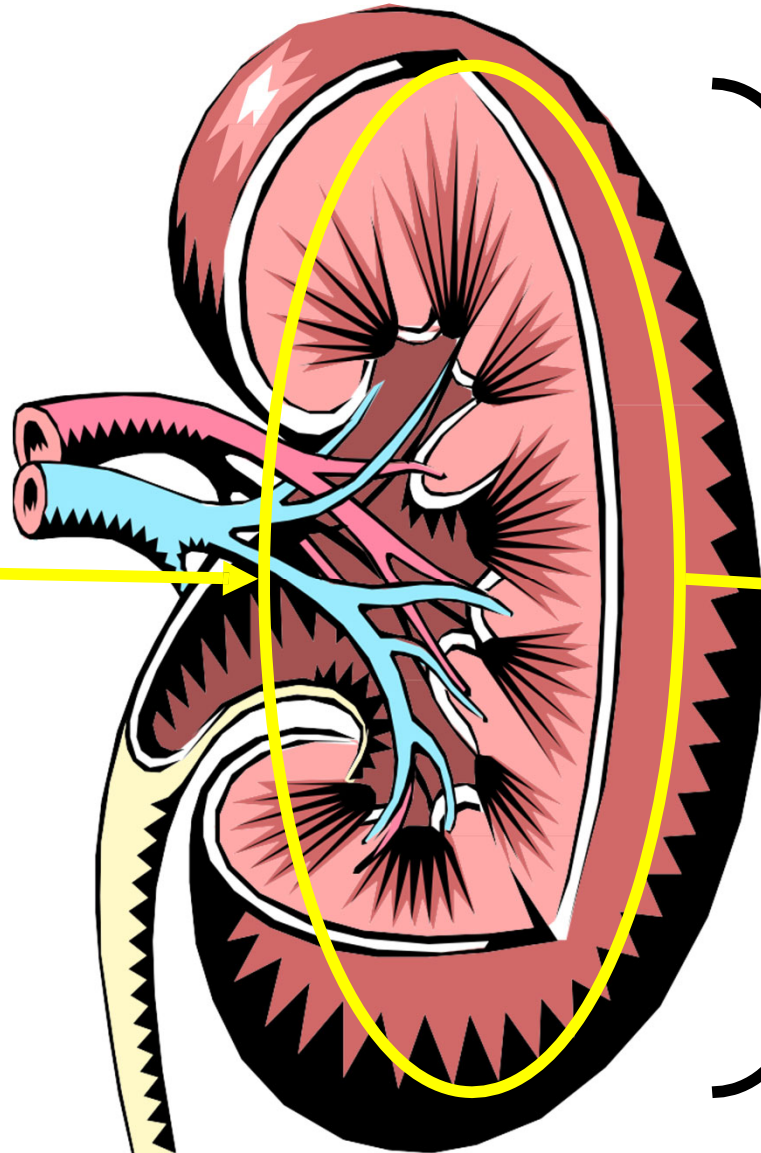
Parenchymal
Structures

↓ GFR

Intrinsic
AKI

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Syndromes of Acute Kidney Injury

Factors that Affect:

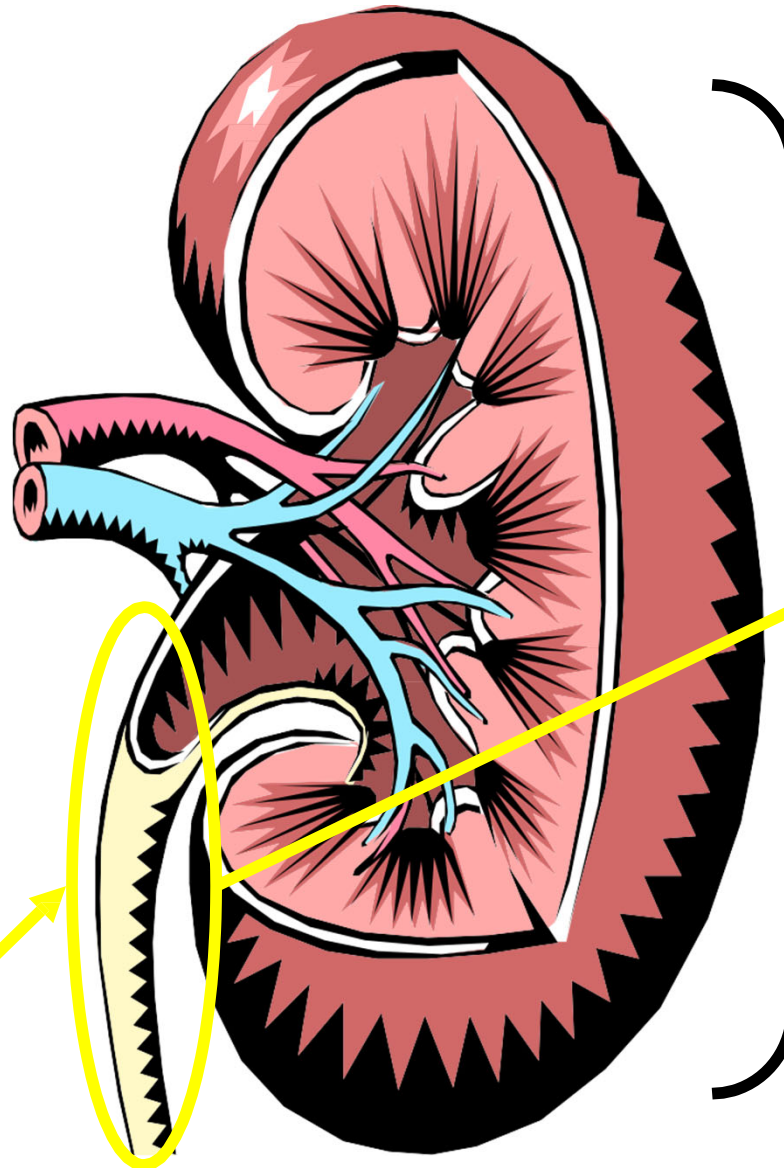
Induce:

Syndrome:

Renal
Perfusion

Parenchymal
Structures

Urine Outlet
Tract



Prerenal
AKI

Intrinsic
AKI

Postrenal
Obstructive

↓ GFR



Acute Kidney Injury

Prerenal AKI	Intrinsic AKI	Postrenal AKI
<p>↓ <u>Intravascular Volume</u> Dehydration / Hemorrhage GI, Cutaneous or Renal losses Third Spacing</p> <p>↓ <u>Effective Blood Volume</u> CHF Cirrhosis Nephrotic Syndrome Sepsis Anesthesia</p> <p><u>Altered Renal Hemodynamics</u> Preglomerular Constriction Postglomerular Vasodilation Medications: ACEI, NSAIDS, CSA Hepatorenal syndrome, Surgery</p> <p><u>Renal Vascular Obstruction</u></p> <p><u>Abdom. Compartment Synd.</u></p>	<p><u>Acute Tubular Necrosis</u> Ischemic: Sepsis Hypotension Nephrotoxic: Drugs Pigments</p> <p><u>Acute Interstitial Nephritis</u> Drug-induced Infection-related Systemic Diseases Malignancy</p> <p><u>Acute Glomerulonephritis</u></p> <p><u>Acute Vascular Syndrome</u> Renal artery dissection Renal artery Throm-Emb Renal vein thrombosis Atheroembolic disease</p>	<p><u>Upper Tract Obstruction</u> Intrinsic Stone Papillary Necrosis Blood Clot TCC Extrinsic Retroperit. Fibrosis Malignancy Ligation Pelvic Mass</p> <p><u>Lower Tract Obstruction</u> Urethral Stricture BPH Prostate Cancer TCC of the bladder Stones: bladder Neurogenic bladder Malpositioned Foley Cath</p>





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Meds: atorvastatin, chlorthalidone, amlodipine, tamsulosin, ASA

Retired military who still works full time @ golf shop

IF his baseline kidney function was SCr of 0.6 and eGFR of 95 and today the SCr and eGFR are 1.4 and 47, what is the cause of kidney dysfunction?

A. Acute kidney injury (AKI). Because GFR of <60 at 47 has been present X1 not for >3 months. So WHAT TYPE of AKI???

~~B. AKI on chronic kidney disease (CKD)~~

~~C. CKD~~



George



80 y/o male with HDL, HTN, BPH, CAD, former smoker
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Presents to **your office** with gross hematuria x 1 week.
Denies dysuria, fever/chills, penile discharge, LE edema, or changes in urinary habits

UA: + heme, negative protein, 3-10 RBC, negative WBC,
25-100 hyaline casts

UACR: (urine albumin/creatinine ratio) <30 WNL

What is George's AKI etiology/diagnosis?

- A. Nephrotic syndrome
- B. Acute interstitial nephritis (AIN)
- C. Acute glomerulonephritis AKI
- D. Post-renal AKI
- E. Acute tubular necrosis (ATN)

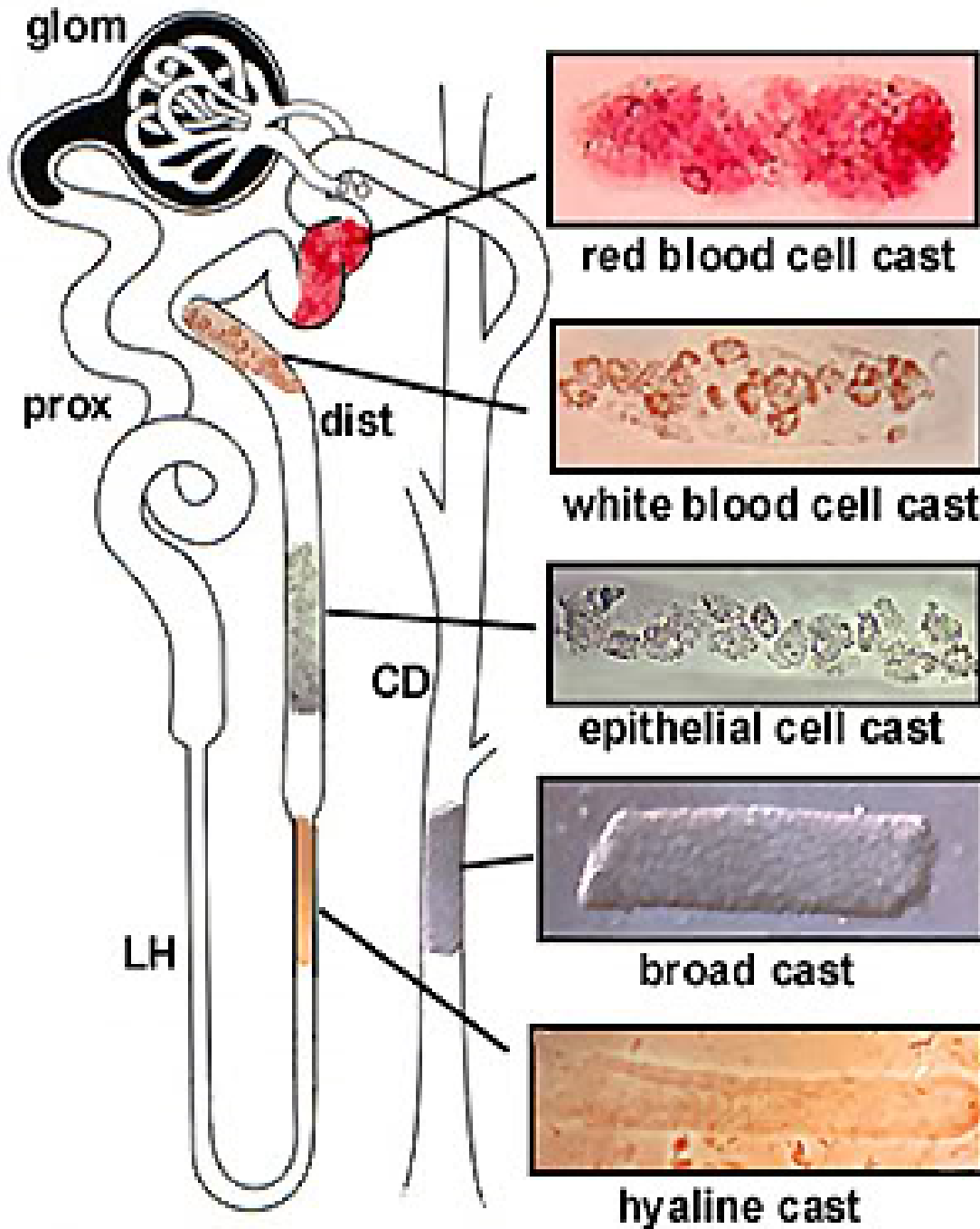


Steps to success in nephrology

1. What is the kidney function? Acute vs chronic
2. What do you see in the urine analysis?
3. How much proteinuria?
4. What do you see on kidney ultrasound?

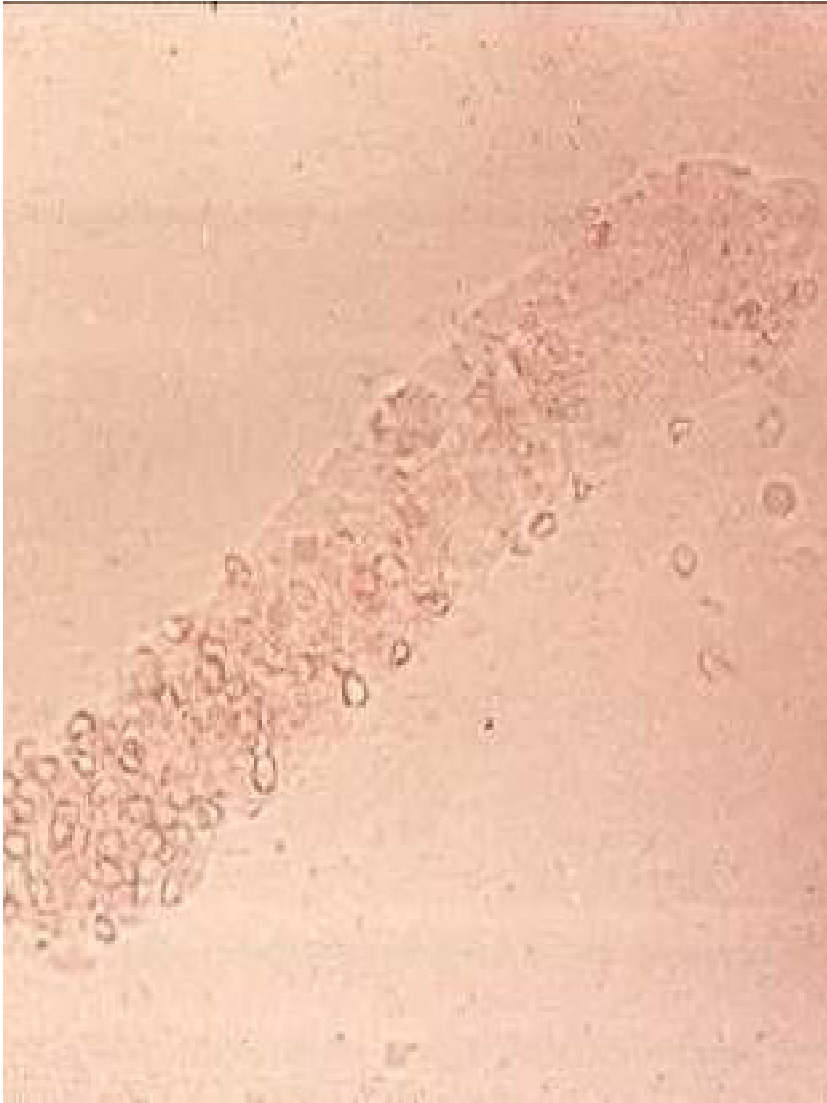


Urinary casts

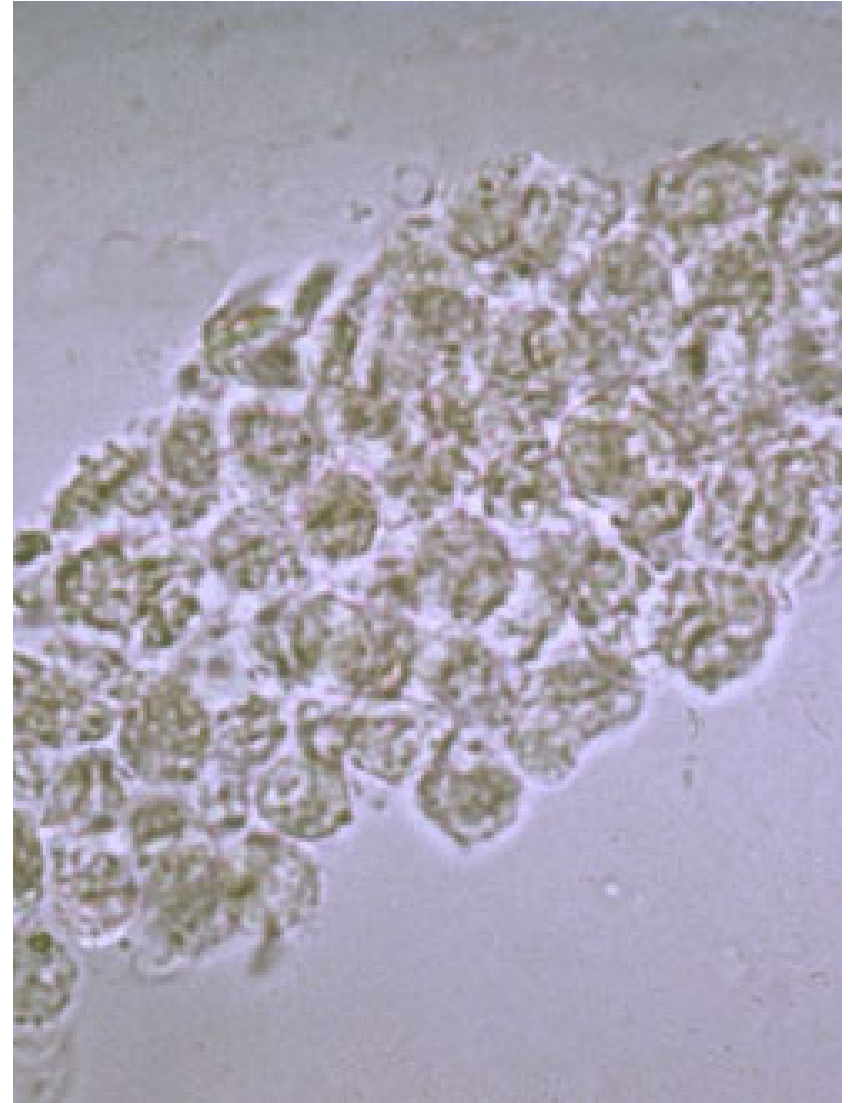


Named based upon shape and origin

1. Hyaline
2. Granular
3. Waxy/Broad
4. RBC
5. WBC
6. Tubular



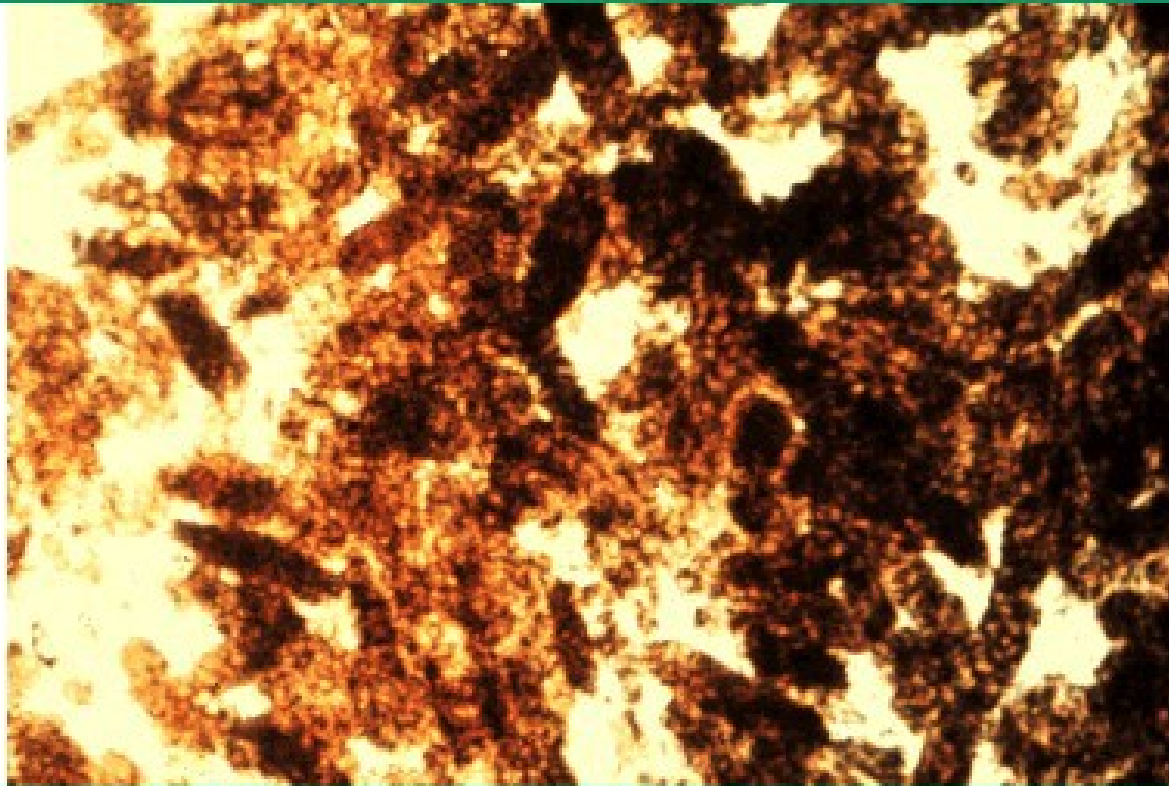
RBC Cast
THINK
GLOMERULONEPHRITIS



WBC Cast
THINK  Fun with Kidneys
INTERSTITIAL NEPHRITIS

Muddy/Granular Brown Casts = ATN

Photomicrograph showing urine sediment with muddy brown granular casts



Pathognomonic for ATN

Hyaline Casts = Hell, if I know NOTHING



Hyaline cast

Not pathognomonic for anything.....



Urine cheat sheet for intrinsic kidney injury

	Cystitis	Pyelonephritis	Acute Allergic Interstitial Nephritis	Acute Glomerulonephritis	Nephrotic Syndrome	Ischemic Tubular Necrosis
Protein	0	0	0	++	++++	0
RBCs	0	0	+	+++	0	0
WBCs	++++	++++	++++	+	0	0
RBC Casts	0	0	0	++++	0	0
WBC Casts	0	+++	+++	0	0	0
Granular Casts	0	+	+	+	0	++++

George



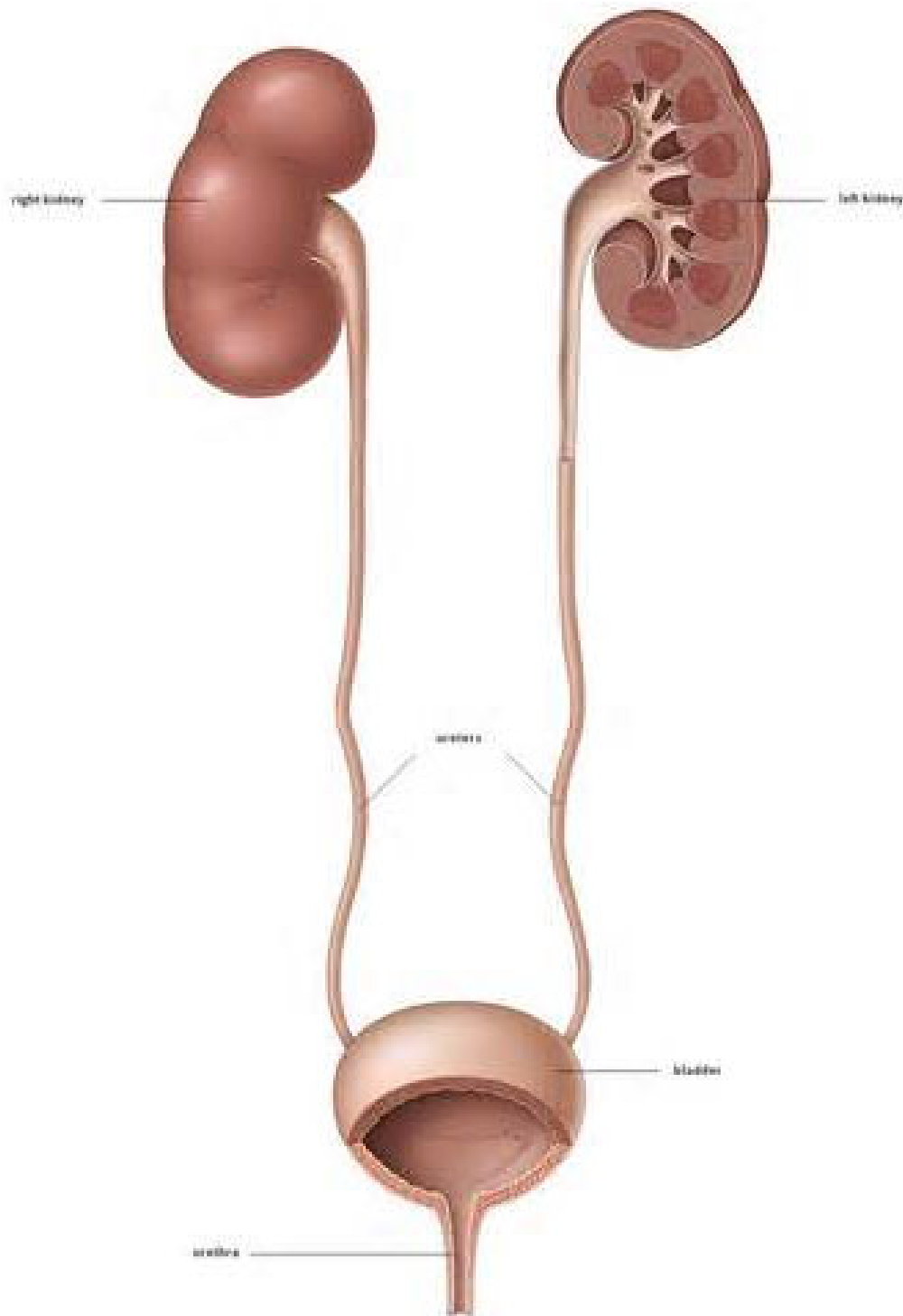
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**UA: + heme, negative protein, 3-10 RBC,
negative WBC, 25-100 hyaline casts
UACR (urine alb/cr ratio) <30 WNL**

What is George's diagnosis?

- ~~A. Nephrotic pattern~~
- ~~B. Acute interstitial nephritis (AIN)~~
- C. Acute glomerulonephritis AKI
- D. Post-renal AKI
- ~~E. Acute tubular necrosis (ATN)~~





Where is the origin of the hematuria?

Causes of hematuria

1. Renal: renal mass, **glomerular** pathologies, structural disease, pyelonephritis, malignant HTN, renal vein thrombosis, AV malformation, papillary necrosis(sickle cell)
2. Ureter: **malignancy**, stone, stricture, fibroepithelial polyp
3. Bladder: **malignancy**, radiation, cystitis
4. Prostate/urethra: BPH, **malignancy**, prostatic procedures, traumatic catheterizations, urethritis, urethral diverticulum



Steps to success in nephrology

1. What is the kidney function?
2. What do you see in the urine analysis?
3. How much proteinuria?
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George



What is George's diagnosis?

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- ~~C. Acute glomerulonephritis AKI~~
- D. Post-renal AKI**
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Kidney sonogram demonstrated moderate **B/L hydronephrosis**. No calculi or renal masses. Noted to have **large mass noted in urinary bladder**.



Sally



78 y/o presents ED with nausea & vomiting for 2 days. Unable to keep food down

No available PMH

PE: 110/60 HR 80 sitting

80/55, HR 100 standing

Labs: BUN 45mg/dL, SCr 1.5mg/dL, FeNa 0.3%

UA: neg heme/protein, 0-1 RBC, neg WBC

25-100 hyaline casts. UACR <30mg/g

What is Sally's diagnosis?

- A. UTI
- B. Acute interstitial nephritis (AIN)
- C. Pre-renal AKI
- D. Post- renal AKI
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No available PMH

PE: 110/60 HR 80 sitting

80/55, HR 100 standing

Labs: BUN 45mg/dL, SCr 1.5mg/dL.

Review of EMR shows SCr 0.9mg/dl in

2018 FeNa 0.3% UA: neg heme/protein, 0-1 RBC, neg WBC, 25-100 hyaline casts. UACR <30mg/g

What is Sally's diagnosis?

- A. UTI
- B. Acute interstitial nephritis (AIN)
- C. Pre-renal AKI
- D. Post- renal AKI
- E. Acute tubular necrosis (ATN)



Steps to success in nephrology

1. What is the kidney function? **Acute** vs chronic
2. **What do you see in the urine analysis?**
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What is Sally's diagnosis?

~~A. UTI~~

~~B. Acute interstitial nephritis (AIN)~~

C. Pre-renal AKI

D. Post-renal AKI

~~E. Acute tubular necrosis (ATN)~~



FeNa (Fractional Excretion of Na)

Remember FeNa is a urine test

	Pre Renal	Intrinsic Renal	
FeNa	<1%	>1%	

Used to help differentiate between intra-renal process or extra-renal

FeNa < 1% ~ prerenal cause, volume depletion

Kidney corrects for low fluid state by reabsorbing Na, therefore functional kidney

FeNa > 1% ~ ATN

Failing kidney...cannot compensate and leaking sodium thus indicates kidney diagnosis

- **Prerenal:** Decreases Kidney Perfusion: Hypovolemia, CHF, Renal Artery Stenosis, Sepsis, etc
- Remember, contrast-induced nephropathy will often look pre-renal!
- **Intrinsic Renal:** ATN



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78 y/o presents ED nausea and vomiting for 2 days. Unable to keep food down

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PE: 110/60 HR 80 sitting

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Renal sonogram was unremarkable without hydronephrosis

What is Sally's diagnosis?

~~A. UTI~~

~~B. Acute interstitial nephritis (AIN)~~

C. Pre-renal AKI

D. Post-renal AKI

~~E. Acute tubular necrosis (ATN)~~



Carrie



85yo with advanced CKD stage 4 (eGFR 28), HTN & uncontrolled type 2 DM with retinopathy who presents to your office with fever, dysuria, malaise and lower abdominal pain x2 days.

BP 125/50, P 105, T 101F

Labs: eGFR 20, SCr 2.5mg/dl, BUN 30mg/dl

UA was negative for leukocyte esterase and positive for nitrite, and positive for glycosuria. 0-2RBC, 25 WBC, 10 WBC casts, UACR <30mg/g

What is Carrie's diagnosis?

- A. Post-renal AKI on CKD
- B. Pre-renal AKI on CKD
- C. Acute tubular necrosis (ATN)
- D. Acute Interstitial nephritis (AIN) on CKD
- E. Acute Interstitial nephritis (AIN)



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BP 125/50, P 105, T 101F

Labs: eGFR 20, SCr 2.5mg/dl, BUN 30mg/dL

**2 months ago, eGFR was 28,
SCr 2.0, BUN 25mg/dl → AKI on CKD**

What is Carrie's diagnosis?

- A. Post-renal AKI on CKD
- B. Pre-renal AKI on CKD
- C. ~~Acute tubular necrosis (ATN)~~
- D. Acute Interstitial nephritis (AIN) on CKD
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Carrie



85yo with advanced CKD stage 4 (eGFR 28), HTN & uncontrolled type 2 DM with retinopathy who presents to your office with fever, dysuria, malaise and lower abdominal pain x2 days. **Pyelonephritis**

Sonogram showed isoechoic kidneys R and L measuring 8.5cm & 8.3cm with evidence of medical renal disease. No hydronephrosis

What is Carrie's diagnosis?

- ~~A. Post-renal AKI on CKD~~
- ~~B. Pre-renal AKI on CKD~~
- ~~C. Acute tubular necrosis (ATN)~~
- D. Acute Interstitial nephritis (AIN) on CKD**
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Mickey

52 y/o male with history of HTN since mid 20s.

No family history, he was adopted

Meds: chlorthalidone and prn ibuprofen

Presents to your office with complaints of intermittent sharp pains in left lower flank and back for past few months. Denies nausea, vomiting, diarrhea, gross hematuria, dysuria, fever/chills.

PE: BP 150/92, P 85. Unremarkable

Labs: Na⁺ 138, K⁺ 5.8, SCr 1.3, BUN 18, eGFR 66

UA: negative heme/protein, negative RBC/WBC, no casts

UACR (urine albumin/creatinine ratio) 200mg/g

What is causing Mickey's kidney dysfunction?

- A. AKI**
- B. CKD**
- C. AKI on CKD**



Steps to success in nephrology

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Trending of kidney function

Differentiating acute vs chronic

Mickey's renal function

<u>SCr</u>	<u>eGFR</u>	<u>UACR</u>	<u>Year</u>
0.77	124	<30	1999
0.99	96	50	2010
1.2	75	70	2012
1.3	66	200	2019





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PE: BP 150/92, P 85. Unremarkable

Labs: Na⁺ 138, K⁺ 4.6, **SCr 1.3**, **BUN 18**, **eGFR 66**

UA: negative heme/protein, negative RBC/WBC, no casts

UACR (urine albumin/creatinine ratio) 200mg/g

What is causing Mickey's kidney dysfunction?

~~A. AKI~~

B. CKD

~~C. AKI on CKD~~



Criteria for CKD

1. Decreased GFR \leq 60ml/min for **> 3 months**
OR
2. Markers of Kidney Damage for **> 3 months**
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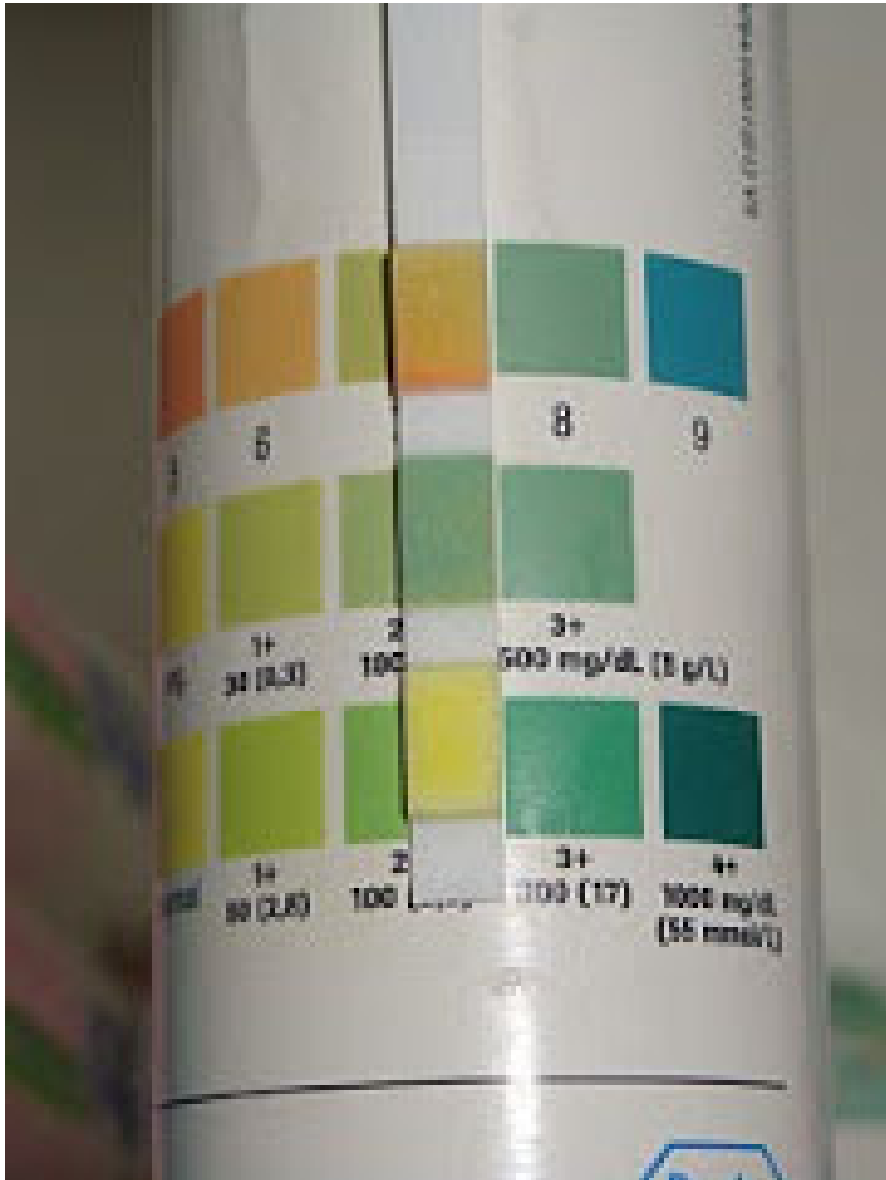
~~A. AKI~~

B. CKD

~~C. AKI on CKD~~



Proteinuria???



negative	0 mg/dL
trace	15-30 mg/dL
1+	30-100 mg/dL
2+	100-300 mg/dL
3+	300-1000 mg/dL
4+	>1000 mg/dL



Quantification of Proteinuria

1. 24 hour total urinary protein $<150\text{mg}$ is
NORMAL

OR

2. Random spot

UPCR: protein/Cr $<150\text{mg/g}$ is WNL

UACR: albumin/Cr $<30\text{mg/g}$ is WNL

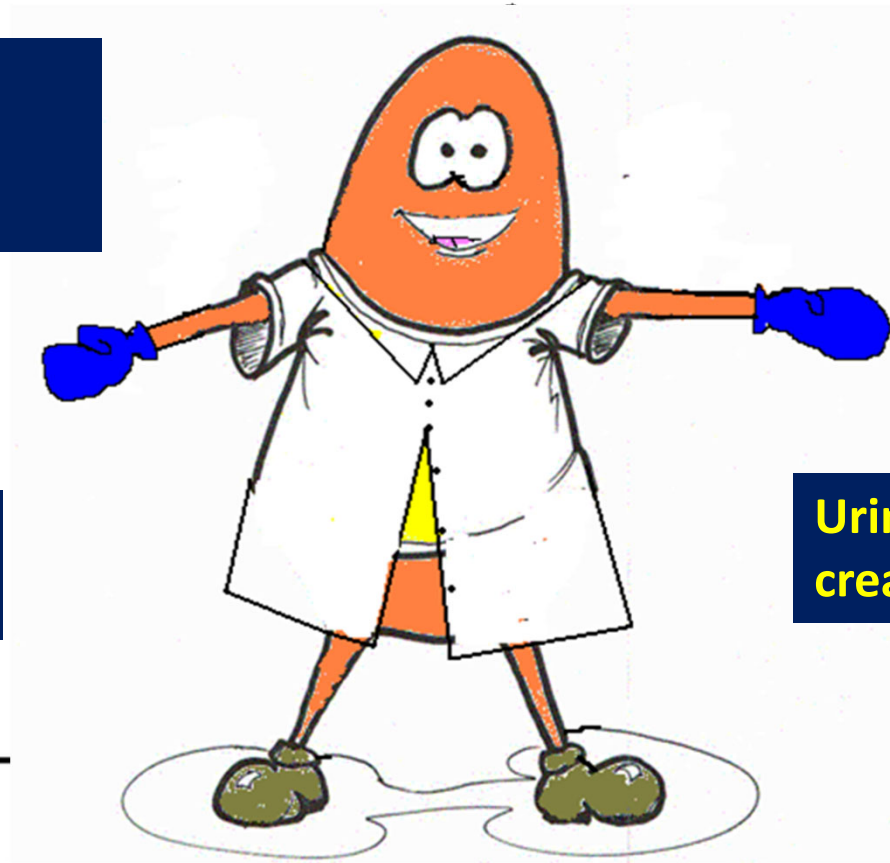
Adjustment for urinary concentration is made
by relating the urine protein concentration to
the urine creatinine concentration



Quantifying **ALBUMINURIA**: Where is the UA dipstick?

Urine protein to
creatinine ratio
(UPCR)

24 hr
for
in



Urine
Or

Urine albumin to
creatinine ratio (UACR)

Special Thanks to Scott and White of Temple TX for
use of their kidney comic



Nephrotic vs Non-nephrotic

- **Nephrotic >3 grams of protein in 24hrs**
- Syndrome associated with
 1. Edema
 2. HTN
 3. Rapid decline in kidney function
 4. Oval Fat bodies
 5. Hypercoag state
 6. Hypertriglyceridemia
- ALL PROTEINURIA is TOXIC to the kidney
- ALL PROTEINURIA has bad prognosis
- More PROTEINURIA worse prognosis
- PROTEINURIA is not just marker of kidney disease

IT IS TOXIC to KIDNEY



Albuminuria Necessary to Stage CKD AND Determine Prognosis

CKD classified based on:

- GFR (G)
- **Albuminuria (A)**

KDIGO 2012

CKD classified based on: ● GFR (G) ● Albuminuria (A) <i>KDIGO 2012</i>				Albuminuria categories		
				Description and range		
				A1	A2	A3
				Normal to mildly increased	Moderately increased	Severely increased
				<30 mg/g <3 mg/mmol	30-299 mg/g 3-29 mg/mmol	≥300 mg/g ≥30 mg/mmol
GFR categories (ml/min/1.73 m ²) Description and range	G1	Normal or high	≥90	Monitor 1	Monitor 1	Refer* 2
	G2	Mildly decreased	60-90	Monitor 1	Monitor 1	Refer* 2
	G3a	Mildly to moderately decreased	45-59	Monitor 1	Monitor 2	Refer 3
	G3b	Moderately to severely decreased	30-44	Monitor 2	Monitor 3	Refer 3
	G4	Severely decreased	15-29	Refer* 3	Refer* 3	Refer 4+
	G5	Kidney failure	<15	Refer 4+	Refer 4+	Refer 4+



Mickey

52 y/o male with history of HTN since mid 20s.

No family history, he was adopted

Meds: chlorthalidone and prn ibuprofen

Presents to your office with complaints of intermittent sharp pains in left lower flank and back for past few months. Denies nausea, vomiting, diarrhea, gross hematuria, dysuria, fever/chills.

PE: BP 150/92, P 85. Unremarkable

Labs: Na⁺ 138, K⁺ 5.8, **SCr 1.3**, **BUN 18**, **eGFR 66**

UA: negative heme/protein, negative RBC/WBC, no casts

UACR (urine albumin/creatinine ratio) 200mg/g

What is causing Mickey's kidney dysfunction?

- A. Nephrolithiasis**
- B. Pyelonephritis**
- C. Chlorthalidone + ibuprofen**
- D. ADPKD, autosomal dominant polycystic kidney dz**



Steps to success in nephrology

1. What is the kidney function? Acute vs chronic
2. What do you see in the urine analysis?
3. How much proteinuria?
4. What do you see on kidney ultrasound?





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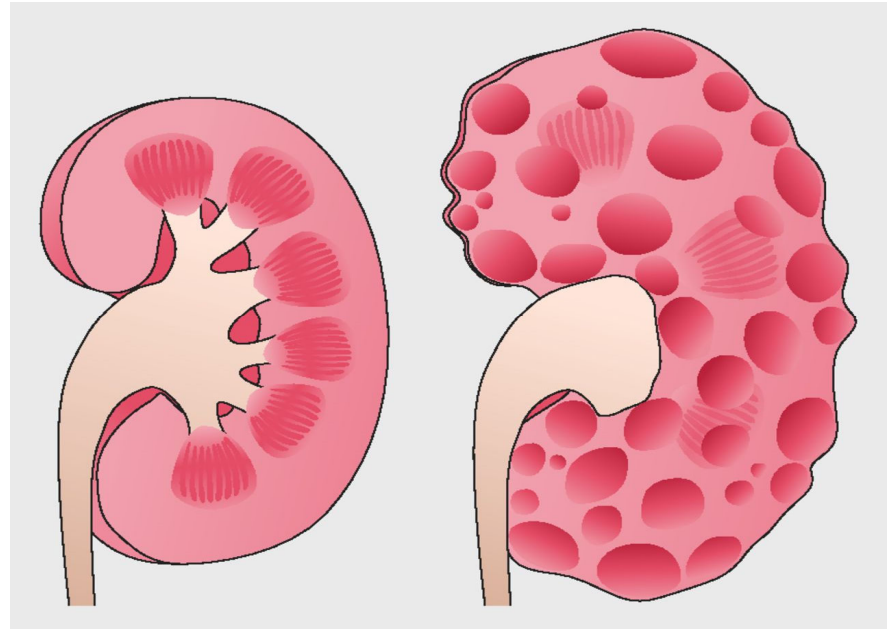
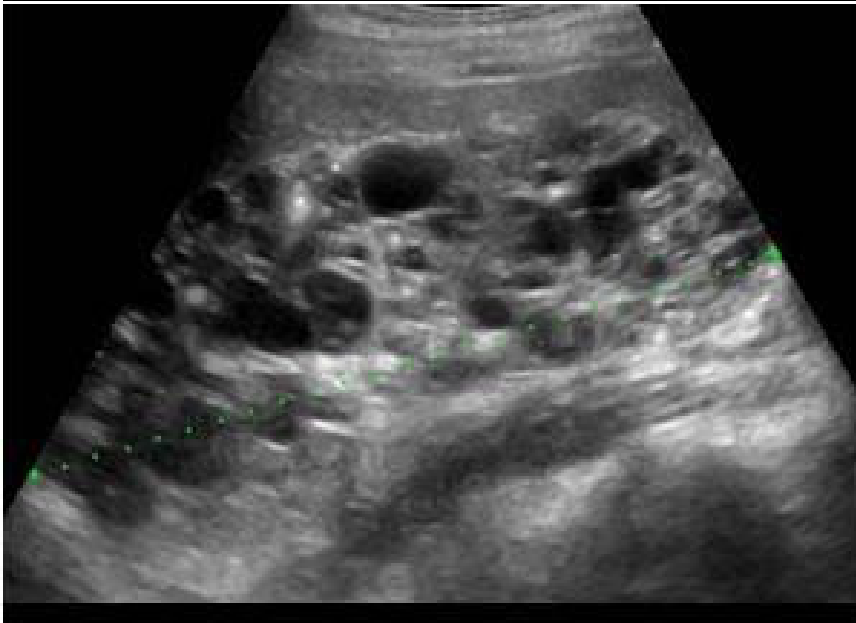
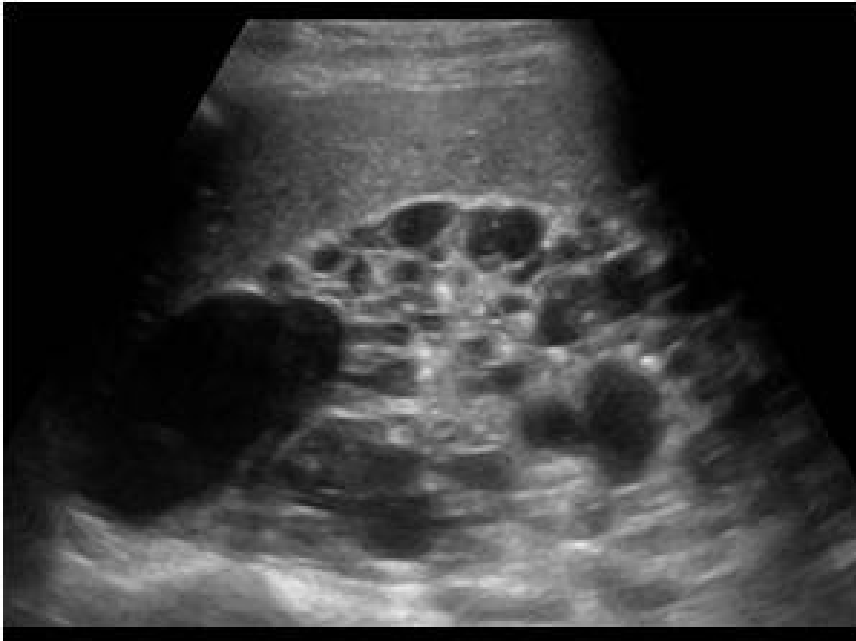
D. ADPKD, autosomal dominant polycystic kidney dz



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IF we knew his family
history, this would
have been an easy
diagnosis

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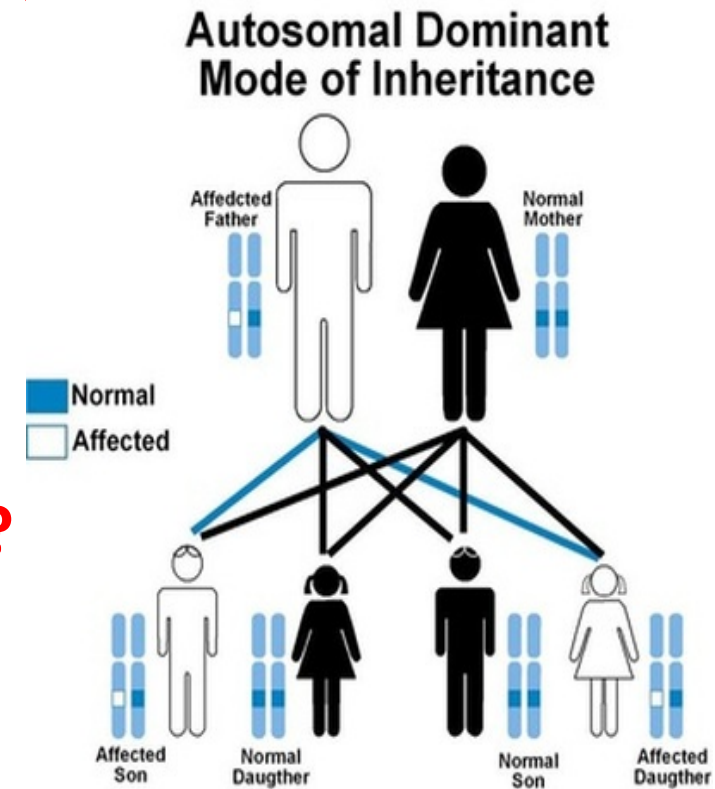
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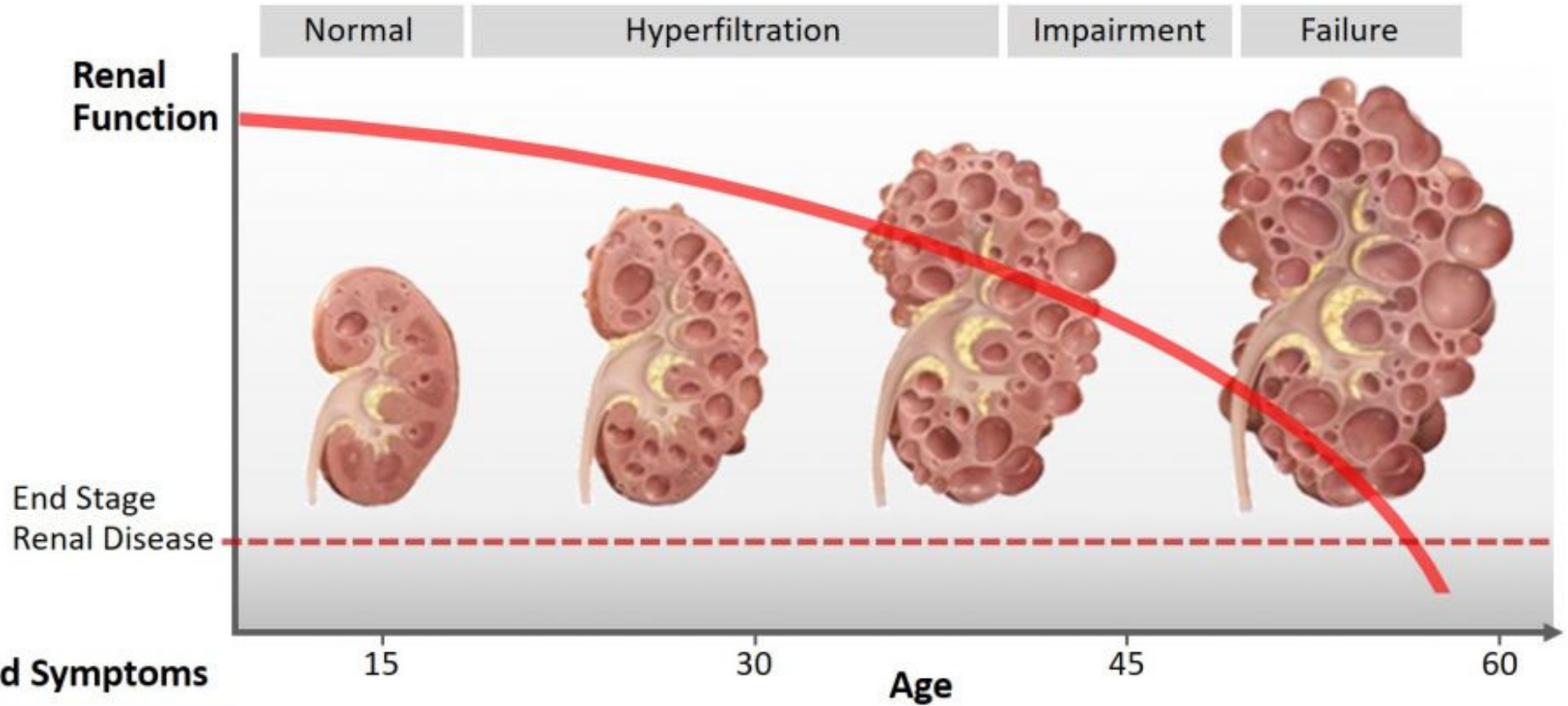
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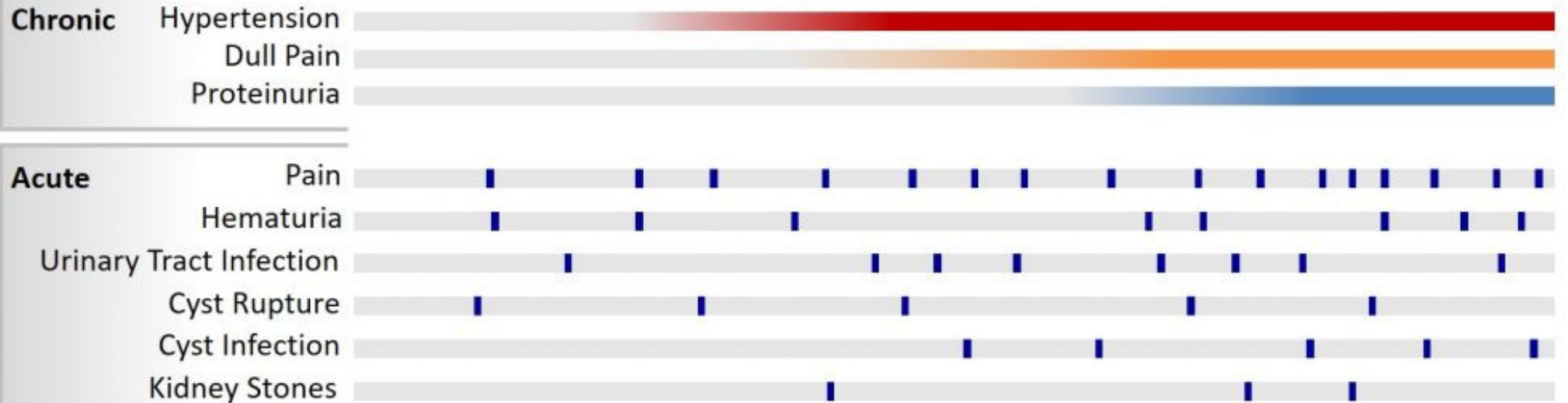
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Typical Disease Progression in ADPKD



Signs and Symptoms





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- A. ACEi usage
- B. Salt substitute
- C. Chlorthalidone
- D. Ibuprofen
- E. CKD



Drug-Induced Hyperkalemia

Mechanisms	Drugs
Impaired RAS function	ACEi/ ARBs, β -blockers Heparin NSAIDs, COX-2 inhibitors
Altered K ⁺ distribution	Insulin antagonists Hypertonic solutions Digoxin, β -blockers
Increased K ⁺ load	Salt substitutes, K ⁺ or herbal supplements PRBC infusions
Reduced K ⁺ excretion	K ⁺ sparing diuretics (MRAs) Calcineurin inhibitors TMP-SMX (Bactrim), pentamidine





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Screening for kidney disease in DM



PMH: Type 2 DM x 15 yrs with retinopathy, obesity, CKD stage 2A3, smoker

Meds: metformin 500mg daily, carvedilol 6.25mg BID

LABS: SCr 1.3mg/dl, eGFR 63ml/min, sK+ 4.0mmol/L, A1C 9.2%, UACR 1500mg/g

PE: BP 145/90, P 80, weight 255lbs, BMI 33, +trace LE edema

Presents for 6 month follow up on DM management.

How would you screen for kidney disease in DM?

1. Evaluate kidney function by looking at eGFR
2. Evaluate kidney function by looking at SCr
3. Quantitate albuminuria by ordering urine albumin/creatinine (UACR)
4. Check urine dipstick for evidence of albuminuria
5. Combination of answers



Screening for CKD in DM

ADA recommendation

- Screen for presence of kidney disease in ALL patients with type 2 diabetes and in type 1 with duration of >5 years
- Annual basis
 1. eGFR = CKD if $<60\text{ml/min}$ for > 3 month
 2. UACR = CKD if $>30\text{mg/g}$ for > 3 month
- If UACR $>300\text{mg/g}$, then check UACR 2x/year to GUIDE THERAPY?



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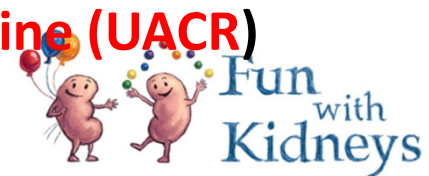
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First steps in treatment of CKD in DM



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1. Refer to dietician for DM education
2. Address lifestyle changes
3. Refer to weight management specialist
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5. All of the above



Diabetic kidney disease risk

537 million

People live with diabetes worldwide

Type 2 diabetes
95%
510 million

Type 1 diabetes
5%
27 million

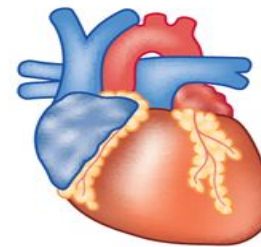
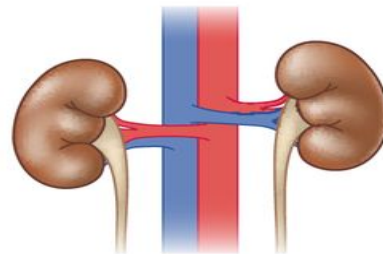


Diabetic kidney disease



Half of chronic kidney disease

Progression to
kidney failure

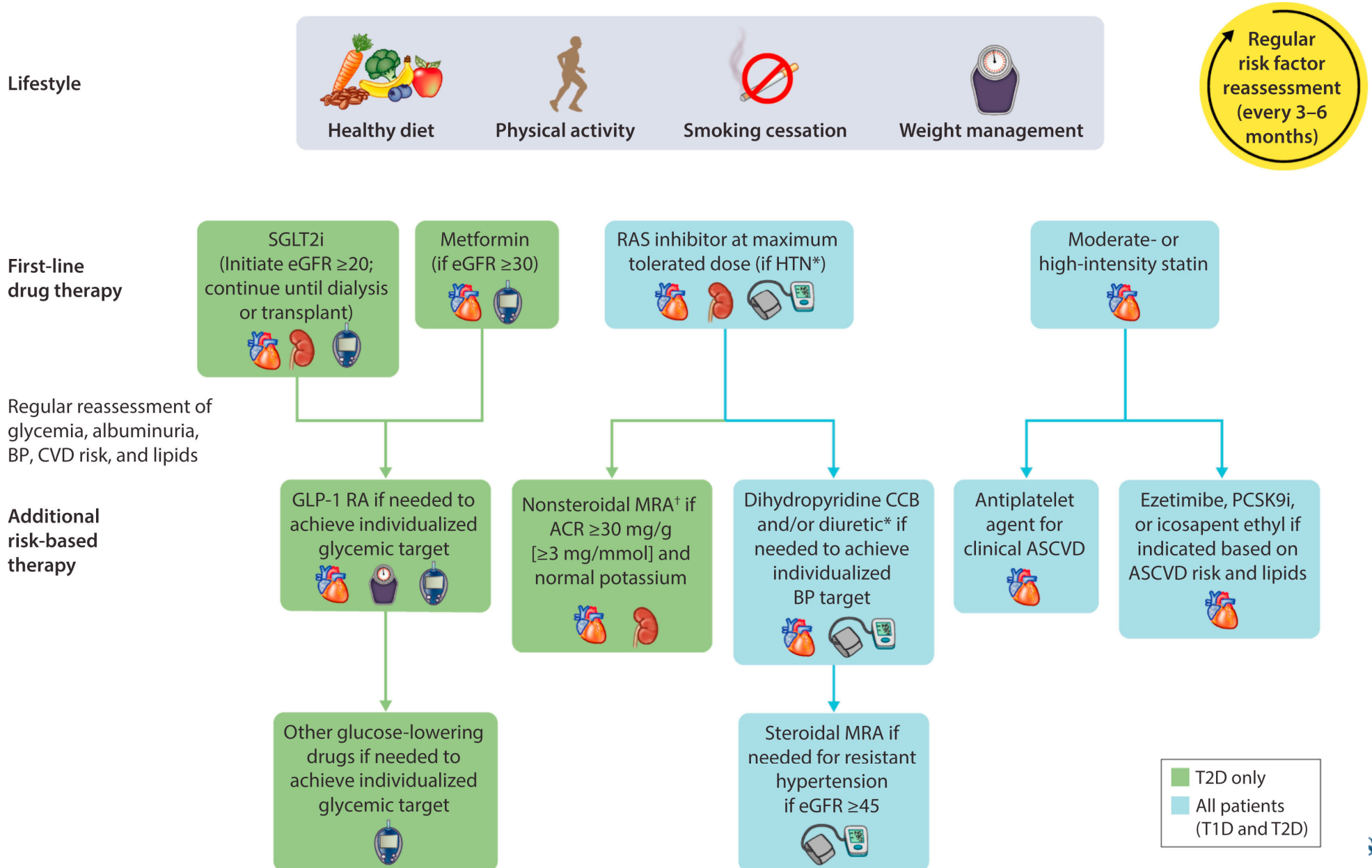


Heart failure,
atherosclerotic
cardiovascular
disease, death

10%

90%

Management of DKD

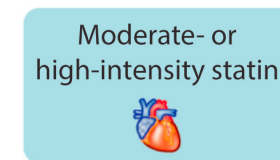
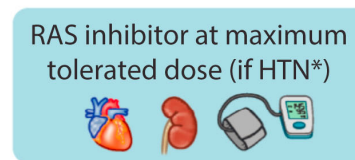
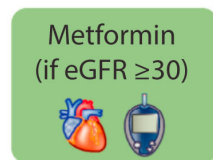
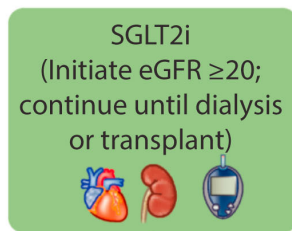


Management of DKD

Lifestyle

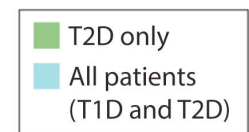
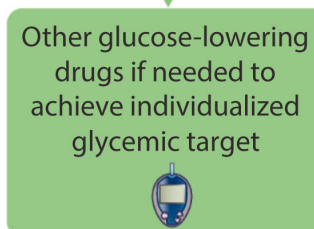
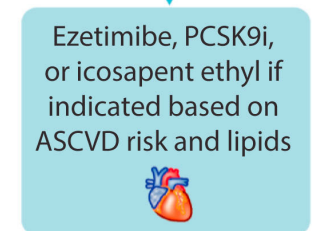
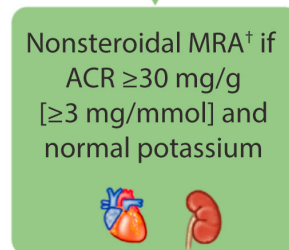
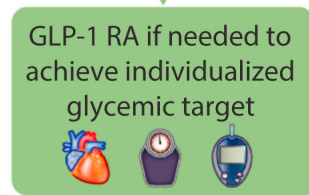


First-line drug therapy



Regular reassessment of
glycemia, albuminuria,
BP, CVD risk, and lipids

Additional
risk-based
therapy



First steps in treatment of CKD in DM



PMH: Type 2 DM x 15 yrs with retinopathy, obesity, CKD stage 2A3, smoker

Meds: metformin 500mg daily, carvedilol 6.25mg BID

LABS: SCr 1.3mg/dl, eGFR 63ml/min, sK+ 4.0mmol/L, A1C 9.2%, UACR 1500mg/g

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Treatment of CKD in DM



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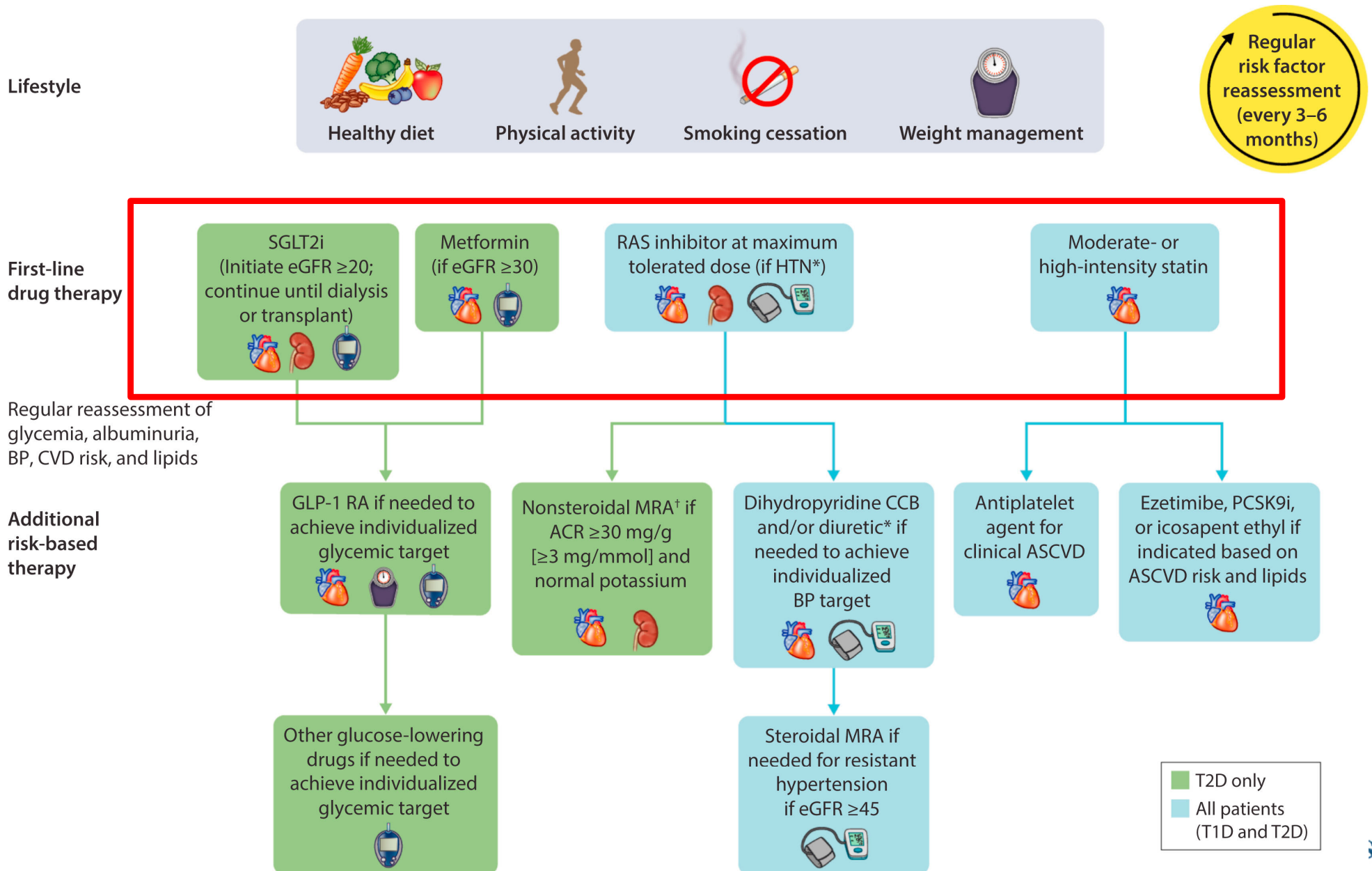
PE: BP 145/90, P 80, weight 265lbs, BMI 33, +trace LE edema

What change would you make for Tx of CKD in DM?

1. Start dapagliflozin 10mg daily
2. Order cholesterol profile
3. Start losartan 25mg daily
4. Consider increasing metformin
5. A combination of multiple answers and if so which answers?



Management of DKD



Treatment of CKD in DM



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Permissive “hypercreatininemia”



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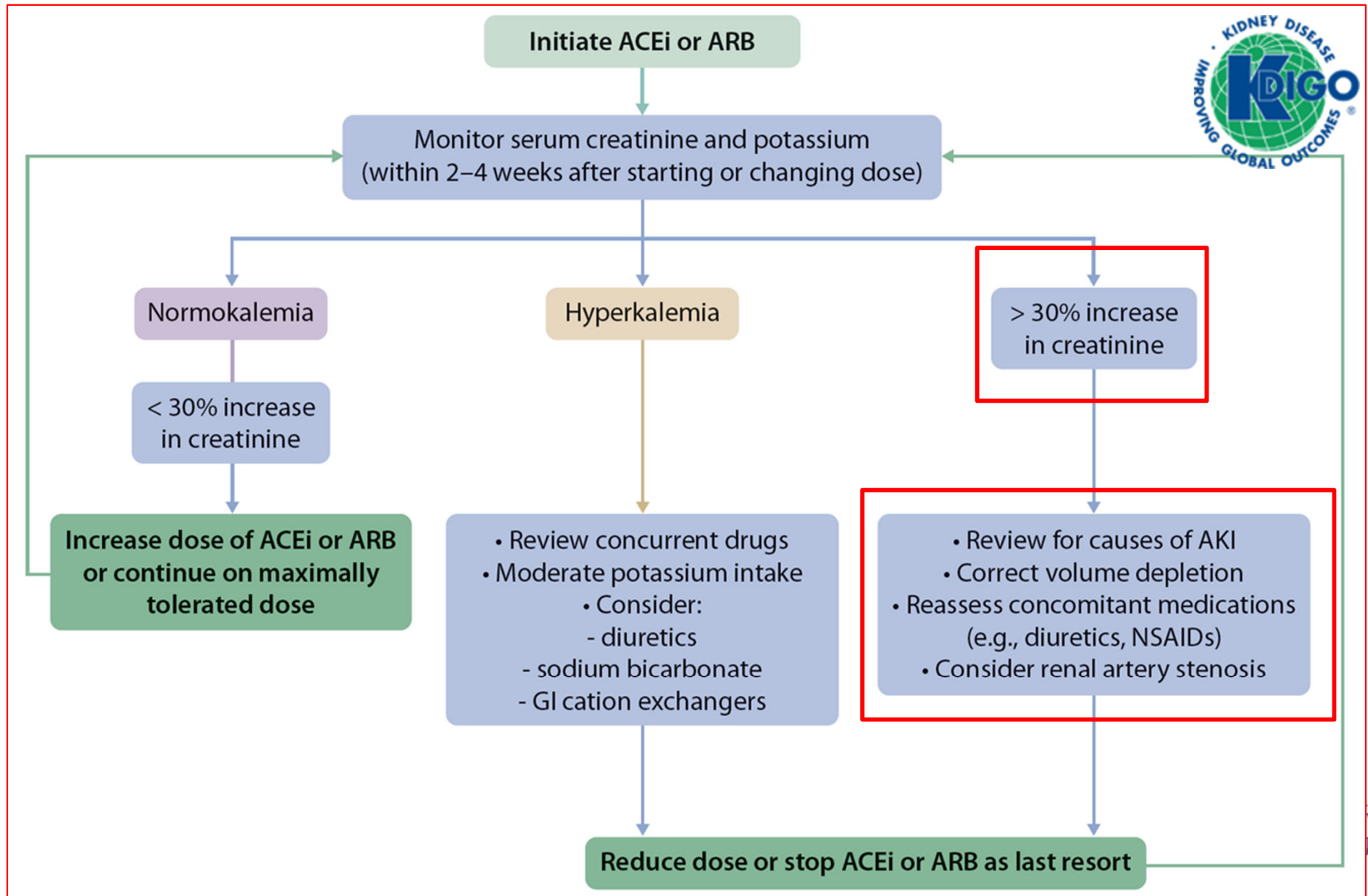
PE: BP 138/85

What would you do next?

1. Look for evidence of pre-renal AKI. Hypotension? Usage of NSAIDs? Vomiting/diarrhea?
2. Decrease losartan to 12.5mg and dapagliflozin to 5mg daily
3. Order renal sonogram with dopplers to ensure no renal artery stenosis
4. Repeat SCr and eGFR in 2 weeks and if stable, continue losartan and dapagliflozin
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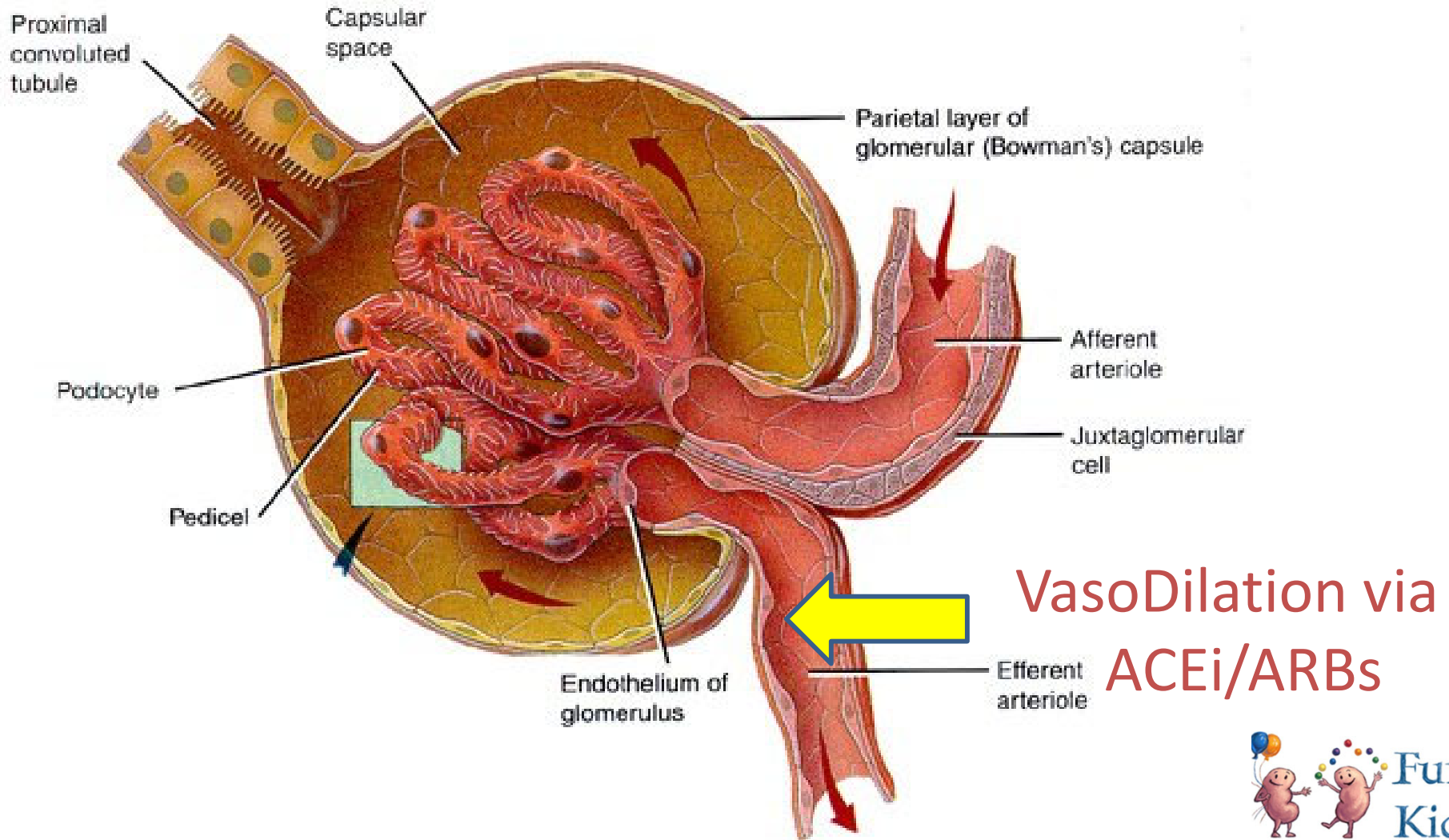


RAS blockade with albuminuria



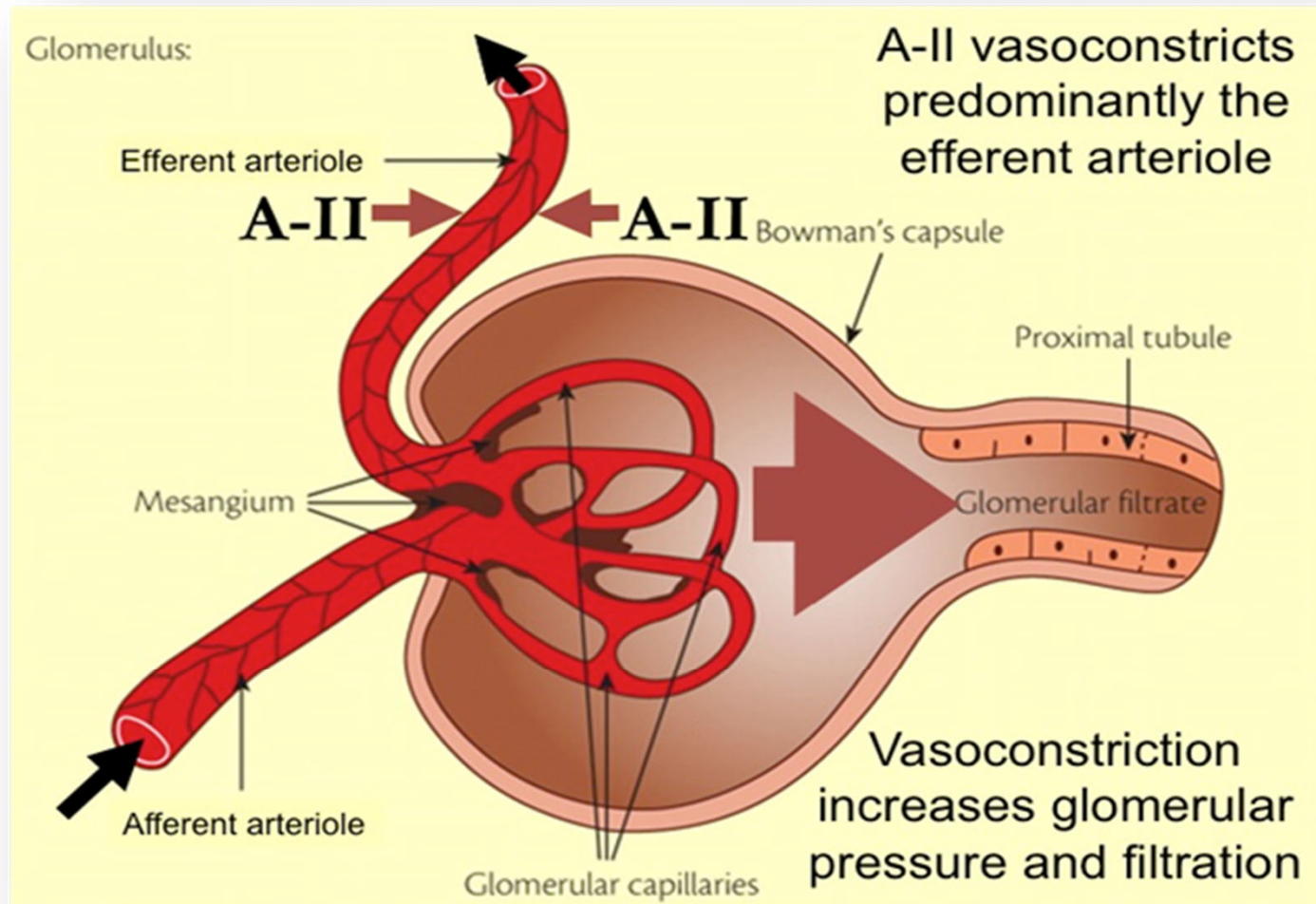
Vaso-dilation efferent arteriole=

GFR + albuminuria



ACEi or ARBs even with advanced CKD

“Permissive hypercreatinemia”



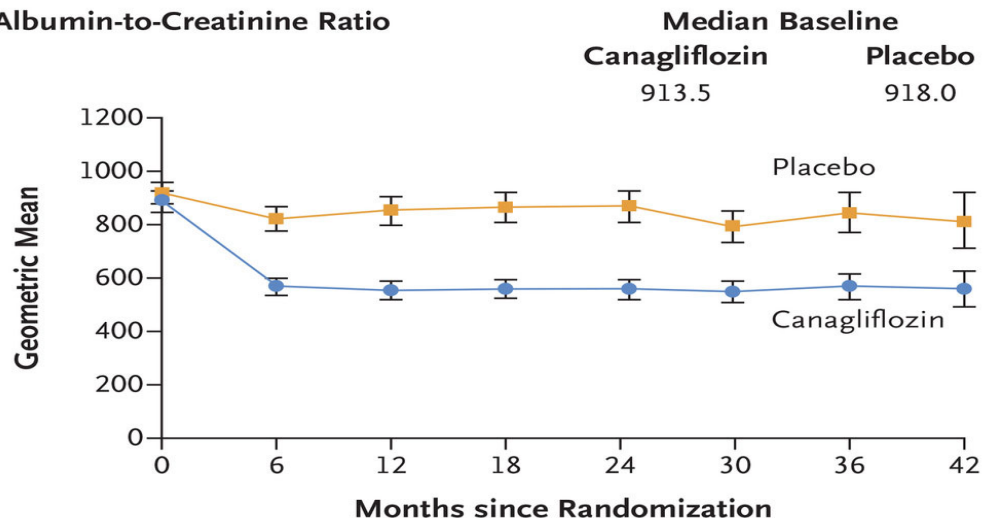
20-30% → 40% bump in SCr is normal. This should be expected. Repeat labs in 2 weeks



CREDESCENCE- SGLT2i

Effects on GFR and albuminuria

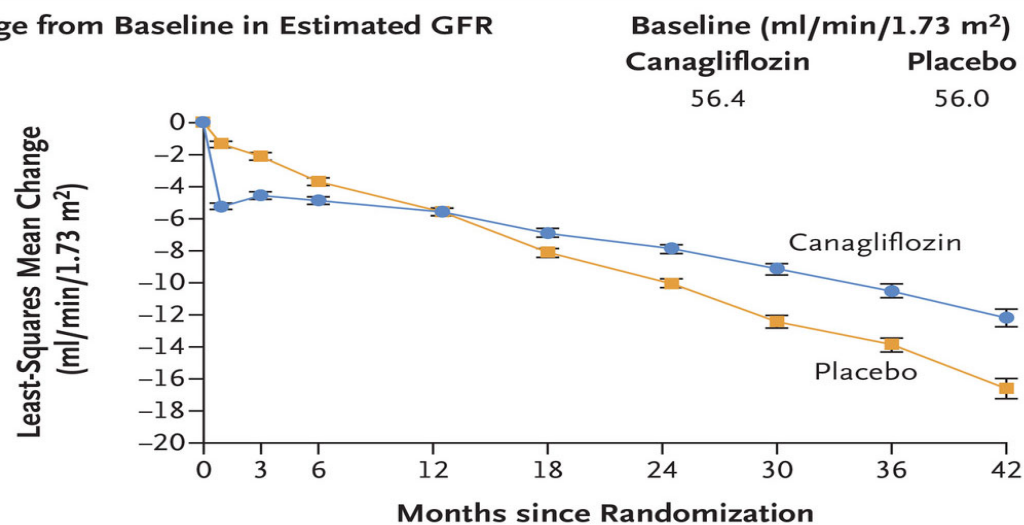
A Urinary Albumin-to-Creatinine Ratio



No. of Patients

Placebo	2113	2061	1986	1865	1714	1158	685	251
Canagliflozin	2114	2070	2019	1917	1819	1245	730	271

B Change from Baseline in Estimated GFR



No. of Patients

Placebo	2178	1985	1882	1720	1536	1006	583	210
Canagliflozin	2179	2005	1919	1782	1648	1116	652	241



SGLT2 Inhibitors

Outcome trials in type 2 DM

- **Reduce risk of major adverse CVD Events**
 - 3-point MACE (myocardial infarction, stroke, CVD death)
 - Heart failure (empagliflozin, canagliflozin, dapagliflozin)
 - CVD death (empagliflozin, dapagliflozin)
- **Decrease severe albuminuria, decline in eGFR, and ESKD.** SGLT2i enhance natriuresis, cause intravascular volume contraction and alter intra-renal hemodynamics, which probably contribute to beneficial effects on blood pressure, body weight and albuminuria



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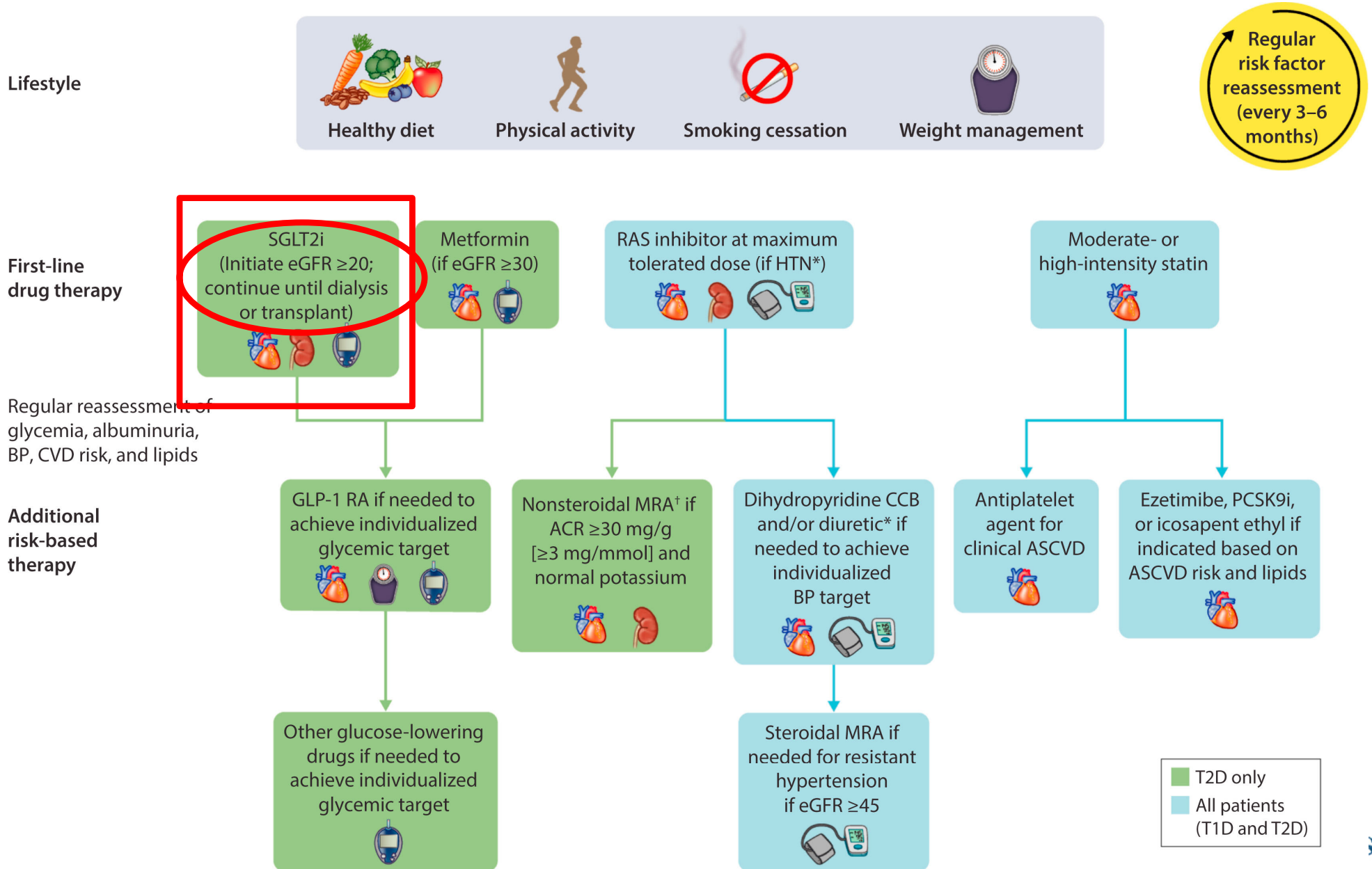
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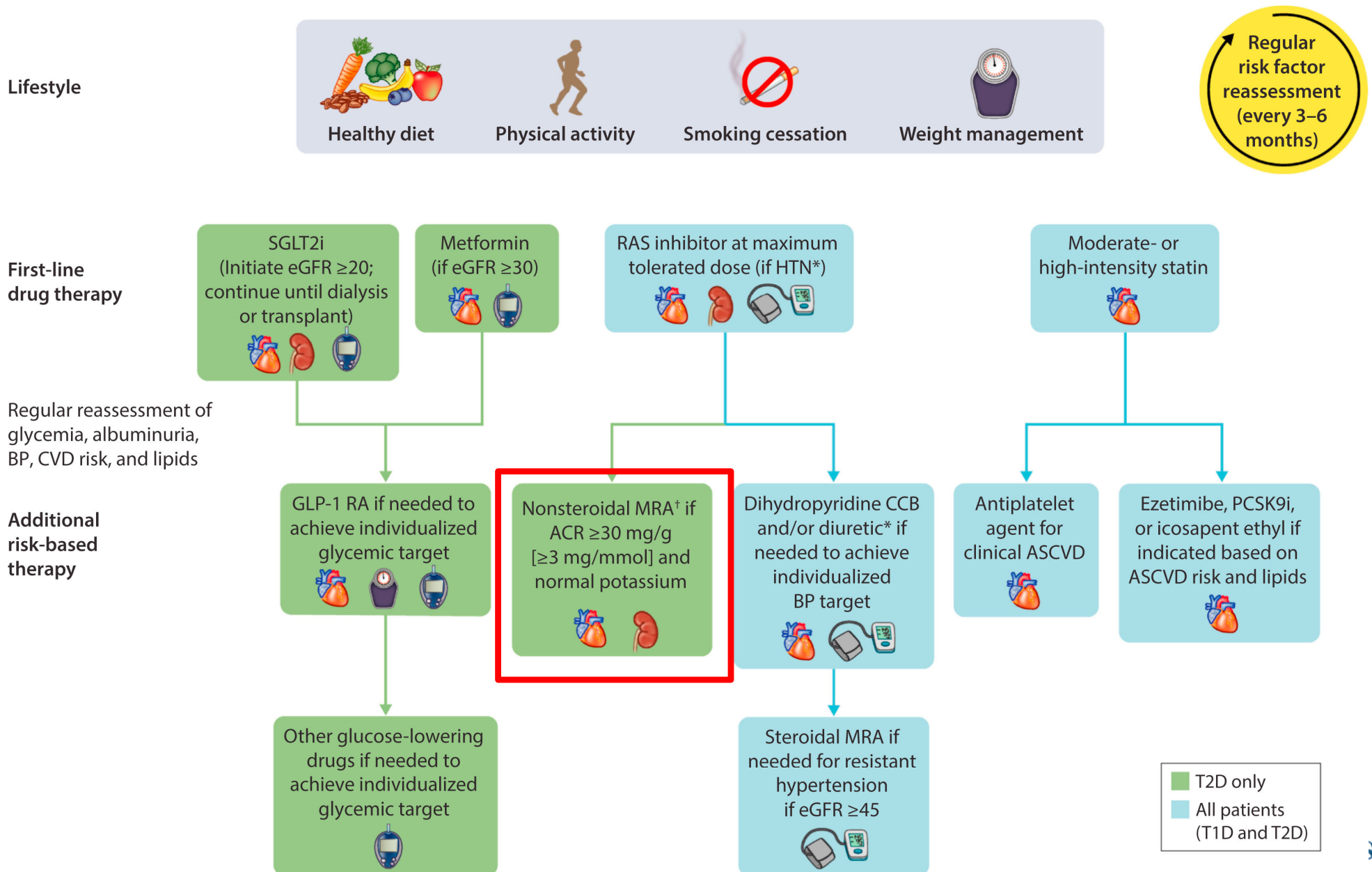
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Management of DKD



Management of DKD



Questions?

Denise.Link@utsouthwestern.edu

