# Ch. 10: Kidney (Renal) Physiology

### **Objectives:**

- 1. Understand renal functions.
- 2. Review anatomy of the urinary system & kidneys.
- **3.** Understand blood flow to kidneys.
- 4. Anatomy & physiology of the nephron.
- 5. Regulation of nephron filtration.
- 6. Kidney disorders.

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## **1. Functions of Renal System**

#### **Regulates:**

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1. Removes metabolic wastes from bloodstream.
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Can live only few weeks (may month) with kidney failure. Patients on kidney dialysis 10 -20 years.

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2. Blood volume - by filtering blood, excreting or reabsorbing water from body as needed (influenced by hormones ______& ____)
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**3. Blood pressure** – by regulating blood volume.

**4. Blood osmolarity** – by controlling reabsorption/excretion of salts (Na+, Cl-, K+, Ca<sup>+2</sup>). Influenced by hormone \_\_\_\_\_

**5.** Blood pH – by controlling reabsorption/excretion of H+ & HCO<sub>3</sub>- in urine.

#### 6. Endocrine functions:

>Calcitrol = increases Ca<sup>+2</sup> absorbed from proximal convol. tubule >Erythropoietin = stimulates RBC production >Renin = secreted by JGA causes \_\_\_\_\_\_

















Bladder Problems contin...

3. Urinary Incontinence – why you can't hold your pee.

<u>A) Urge incontinence</u> = bladder dysfunction. After strong urge to urinate might leak a little (or a lot) urine.



> 1 / 3 people – more often in women
 > common in women w/age & after pregnancy
 > Tx













# Getting a urine sample: Voided sample = collected from normal urination (through urethra) in sample cup. - Can contain sloughed urethral cells and possible bacteria from lower urinary tract. Catheterization = insert (Foley) catheter up urethra into bladder. Bladder Uterus/womb Figure 1 – Female catheter Figure 2 – Male catheter

# Why catheterize the bladder?

## Catheterization

- 3 reasons for catheterization:
- To obtain a sterile urine sample for analysis
- To relieve urinary retention
- To instill medicine into the bladder, after the bladder is emptied
- For urine sample: Quick Cath, In & Out Cath
- For incontinence: Foley Catheter



## **3. REVIEW Blood Supply of Kidneys and nephrons:**

**Renal artery** – brings arterial blood to kidneys to be filtered. – BP in renal artery sensed by the JGA

#### Afferent arteriole = arterial blood enters the glomerulus of the nephrons Plasma, ions, glucose, small proteins, and other substances get filtered through glomerular pores. "Filtrate" then enters PCT.

**Efferent arteriole** = arterial blood leaves the glomerulus of the nephrons (RBCs, WBCs, platelets, and large molecules do not make it through glomerular pores.)

**Peritubular capillaries** = capillaries that surround nephron and receive reabsorbed substances, from filtrate, which return to the bloodstream. OR secrete substances into the filtrate to be removed in the urine.

#### Abbreviations for nephron tubules:

PCT = proximal convoluted tubule Loop = Loop of Henle DCT = distal convoluted tubule





## Review ٠ Functions of renal system - blood volume, pressure, osmolarity endocrine functions • Urinary problems - infections incontinence Anatomy of renal system ٠ - Bladder detrusor muscle - Urethra and sphincters Kidney anatomy - blood supply (renal artery, afferent & efferent arteriole, peritubular capillaries. 20

## 4. Physiology of the Nephron & Blood Filtration

#### Renal Corpuscle = Glomerulus + Bowman's capsule:

A) Glomerulus = receives arterial blood from afferent arteriole, and filters it.
 > has small pores (slits) to allow fluids, ions, glucose, small proteins through.
 > do not allow large molecules or cells (RBCs, WBCs, platelets) through.

**B)** Bowman's capsule = capsule surrounding the glomerulus. Receive the filtrate into the nephron tubule.



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**Glomerular Filtration Rate (GFR)** = volume of filtrate produced by both kidneys per minute. (ml/min)

- Kidneys Filter:

> average of 5.5 L blood every 40 min (entire blood volume!)

- Females = ~ 115 ml/min
- Males = ~ 125 ml/min

\*\*\*GFR is constant for systolic arterial blood pressure (SBP) between 80 – 160 mmHg due to "intrinsic regulation".

The only time GFR changes is when SBP drops below 80 mmHg or goes above 160 mmHg – then it's an "<u>emergency</u>" or extrinsic regulation.









## After Glomerulus & Bowman's capsule → <u>4 Types of Tubules in Nephron:</u> 1. Proximal convoluted tubule > First tubule after glomerulus > Reabsorbs majority of substances from filtrate automatically, regardless of hydration or hormones. 2. Henle's loop > where urine concentrated by Counter-Current multiplication system 3. Distal convoluted tubule > last tubule > where aldosterone has effect on salt 4. Collecting duct – what leaves here is urine.



	Bowman's				
Loop of Henle :	Proximal Convoluted Tubule (PCT) A.a.				
Descending Loop					
> From PCT down to bend	Cortex Culturation				
> Permeable to water but not salt!					
> Where additional 20% of filtrate autor into bloodstream.	matically reabsorbed Descending Loop Henle				
> At bend in loop – between PCT and o of filtrate has been automatically reabs	descending loop ~ 85%				
Things reabsorbed from filtrate:	Things secreted into filtrate				
WATER					







- Addison's Disease Insufficient Aldosterone
- Conn's Syndrome (Hyper-aldosteronism)





### Review Intrinsic regulation of GFR - GFR steady with minor BP fluctuations by afferent arteriole vasodilation / vasoconstriction endocrine functions Extrinsic regulation (medulla) of GFR - If BP $\downarrow$ sympathetic stim $\downarrow$ GFR causing $\downarrow$ urine output and $\uparrow$ blood volume and BP. – If BP $\uparrow$ parasympath. stim $\uparrow$ GFR causing $\uparrow$ urine output and $\downarrow$ blood volume and BP. Structure of nephron - Glomerulus – PCT (what is reabsorbed & secreted?) – Loop of Henle : descending (what is reabsorbed & secreted?) ascending (what is reabsorbed & secreted?) – DCT (what is reabsorbed & secreted? What hormone influences?) - CD (what is reabsorbed? What hormone influences?) 36



# 6. Kidney Disorders

Urinary Stones ("urolithiasis") – frequency 1 / 10 people

Calculi can block renal calyx, ureter, and in males even urethra).

#### <u>Result</u>

> is buildup of fluid pressure (hydronephrosis) within kidneys, causes pressure necrosis.

- > buildup of toxins in bloodstream, causes organs to shut down.





## Urinary Stones ("urolithiasis")

#### Hydronephrosis =

- > can cause pressure necrosis.
- > buildup of toxins in bloodstream, causes organs to shut down.
- > Can also be caused by infection, enlarged prostate (males).



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# **CLINICAL APPLICATIONS**

**Urinary Kidney stones** are composed of crystals (of calcium oxalate, calcium phosphate, and other substances) and proteins that grow until they break loose and pass into the urine collection system. When a stone breaks loose and passes into a ureter, it produces steadily increasing pain, which can become so intense that the patient requires narcotic drugs. The calcium and other substances in kidney stones are normally present in urine, but they become supersaturated and crystalize to form stones for reasons not currently understood. The stones may be removed surgically or broken up by a noninvasive procedure called *shock-wave lithotripsy*.



# Polycystic kidney disease – 1 / 500,000 people USA

#### Result:

Similar to kidney stones. Cysts cause pressure within kidneys. Kidneys become enlarged and pressure causes damage.



# **Chronic Kidney Disease (CKD)**

> 1 / 7 people, USA

> Majority (9/10 people) with CKD don't even know they have it.

> Metabolic blood panel tests look at several factors to asses kidney function (BUN, Creatine, and BUN/Creatine ratio)

Ordered Items Basic Metabolic Panel (8)				
TESTS	RESULT	FLAG	UNITS RE	FERENCE INTERVAL
Basic Metabolic Panel (8)				
Glucose	101	High	mg/dL	65-99
BUN	27	High	mg/dL	6-20
Creatinine	1.01	High	mg/dL	0.57-1.00
eGFR If NonAfricn Am	73		mL/min/1.73	>59
eGFR If Africn Am	85		mL/min/1.73	>59
BUN/Creatinine Ratio	27	High		9-23
Sodium	131	Low	mmol/L	134-144
Potassium	3.2	Low	mmol/L	3.5-5.2
Chloride	109	High	mmol/L	96-106
Carbon Dioxide, Total	34	High	mmol/L	20-29
Calcium	11.2	High	mg/dL	8.7-10.2

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The creatinine clearance test **compares the creatinine level in urine with the creatinine level in blood**. This gives **an estimate of the glomerular filtration rate (GFR).** GFR is a measure of how well the kidneys are working.



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5 Stages of Kidney Disease			
Do NOT memorize!	Kidney Function/GFR	Description	
Stage 1	> 90%	Normal or High Function	
Stage 2	60-89%	Mildly Decreased Function	
Stage 3	30-59%	Mild to Moderately Decreased Function	
Stage 4	15-29%	Severely Decreased Function	
Stage 5	< 15%	Kidney Failure	

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# **Dialysis**

= process of removing excess water, solutes and toxins from the blood in those whose kidneys can't perform anymore. This is also referred to as **renal replacement therap**y.



## **CLINICAL APPLICATIONS**

The term *dialysis* refers to the separation of molecules using their ability to diffuse across an artificial semipermeable membrane (chapter 3). This principle is used in the "artificial kidney" machine for **hemodialysis**. Like the walls of the glomerular capillaries, the artificial semipermeable membrane allows water and dissolved waste molecules (such as urea) to easily diffuse through the membrane pores, whereas plasma proteins are excluded by their larger size. However, unlike the tubules, the artificial membrane can't reabsorb Na<sup>+</sup>, K<sup>+</sup>, glucose, and other molecules needed in the blood. These substances are therefore included in the fluid around the dialysis membrane, so that there is no concentration gradient to cause their net diffusion out of the blood. Sometimes a patient's own peritoneal membranes (which line the abdominal cavity) are used as the dialysis membrane, in a technique called **continuous ambulatory peritoneal dialysis**.

Click <u>HERE</u> for YouTube video explaining dialysis methods 3 min 30 sec

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# **Diuretics**

#### 1. Carbonic anhydrase inhibitors (Acetazolamide)

> Decreases salt reabsorption at PCT. BUT prescribed more for glaucoma patients than patients with hypertension.

#### 2. Loop diuretics (e.g. furosemide or "Lasix")

> Decreases salt reabsorption at ascending loop of Henle. (which will decrease water reabsorption at descending loop!)

Most powerful diuretic, BUT use can lead to loss of other ions (K+, Ca+2, Cl-, and Mg)

#### 3. Thiazides (hydrochlorothiazide)

> decreases salt reabsorption at DCT (BUT can also lead to K+ loss)



# PHYSIOLOGY IN HEALTH AND DISEASE Not required reading

A diuretic is a substance that increases urine volume. *Water* is the most common diuretic, acting to dilute the plasma (lower its osmolarity) and thereby reduce the stimulation of osmoreceptors in the hypothalamus. This lowers the secretion of ADH from the posterior pituitary, which reduces the permeability of the collecting ducts to water and causes *diuresis* (increased water excretion in the urine).

**Osmotic diuretics** are extra solutes in the tubular fluid. These increase the osmolarity of the fluid within the collecting ducts, so that the osmotic gradient (difference in concentration) between the tubular fluid and the interstitial fluid of the renal medulla is reduced. As a result, less water can be drawn out of the collecting ducts by osmosis, leaving more to be excreted in the urine. Glucose is an example of an endogenous molecule that can become an osmotic diuretic, if a person is hyperglycemic and the renal plasma threshold for glucose is exceeded. Because of this, a person with uncontrolled diabetes mellitus who "spills glucose" in the urine has polyuria (literally, "many urines") and can become dehydrated. Similarly, excessive production of ketone bodies (which can cause ketoacidosis; chapter 12) in uncontrolled type 1 diabetes mellitus results in ketonuria, and the extra ketone bodies in the tubular filtrate have an osmotic diuretic effect. A person on a strict weight-reducing diet, who has a rapid breakdown of fat and thus a high plasma level of ketone bodies (ketosis), can also have ketonuria. The resulting osmotic diuresis promotes dehydration, which is part of the reason dieters are advised to drink lots of water. Mannitol is an exogenous substance sometimes used clinically as an osmotic diuretic.

The most powerful clinical diurctics are the **loop diurctics**, including *furosemide* (*Lasix*). These inhibit as much as 25%

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of the salt transport out of the ascending limbs of the loops of Henle. Because of this, the interstitial fluid of the renal medulla is less concentrated (hypertonic), producing less of an osmotic gradient to draw water out of the collecting ducts. The thiazide **diuretics** (such as *hydrochlorothiazide*) inhibit up to 8% of the salt and water reabsorption by inhibiting Na<sup>+</sup> transport in the last part of the ascending limb and first part of the distal tubule, thereby reducing the osmotic gradient for water reabsorption. Although these are effective and commonly used diuretics, Lasix and hydrochlorothiazide have an undesirable side effect: they promote the excretion of  $K^+$  in the urine, which lowers the plasma K<sup>+</sup> concentration (hypokalemia). Hypokalemia can cause neuromuscular disorders and ECG abnormalities. Because of this, people taking Lasix and hydrochlorothiazide should get their blood K<sup>+</sup> concentrations measured periodically, and must often take potassium supplements (in the form of KCl).

The hypokalemia in people taking Lasix or hydrochlo-Not required reading rothiazide is caused by an increase in aldosterone-stimulated secretion of K<sup>+</sup> into the cortical collecting ducts. Because of this, some medications for the treatment of hypertension (high blood pressure; chapter 10) combine hydrochlorothiazide with one of the potassium-sparing diurctics. Spironolactone (such as Aldactone) diuretics block aldosterone action by competing for the aldosterone receptor proteins in the cells of the cortical collecting ducts. Triamterene (Dyrenium) is a potassium-sparing diurctic that acts more directly to block Na<sup>+</sup> reabsorption and K<sup>+</sup> secretion in the cortical collecting ducts. The diuretic actions of hydrochlorothiazide combined with the weaker diuretic but potassium-sparing actions of these drugs lower the blood volume, and thus the blood pressure, of people with hypertension. [Interesting, but too much info (too complex For this course

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# Review

# 3 ways the body regulates blood volume & BP (all involve kidney function!)

- Baroreceptors in heart and medulla influenced by BP and change GFR.
- Hypothalamic ADH (influence by blood osmolarity & change water reabsorption)
- Renin-angiotensin-aldosterone system (influenced by BP and change salt reabsorption)

#### **Kidney Disorders**

- Urinary stones (urolithiasis)
- Polycystic kidney disease
- Chronic kidney disease
- Testing for kidney disease (BUN, Creatinine)
- Kidney failure
- Dialysis
- Diuretics

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