

# Renal Physiology Lect-3

Dr. Ebaa M Alzayadneh,PhD Physiology Department The University of Jordan

Neurohumoral

• Local (Intrinsic)

## 1. Sympathetic Nervous System /catecholamines $\uparrow R_A + \uparrow R_E \longrightarrow \downarrow GFR + \downarrow RBF$

e.g. severe hemorrhage

2. Angiotensin II  $R_E \longrightarrow GFR + \downarrow RBF$ (prevents a decrease in GFR)

e.g. low sodium diet, volume depletion

3. Prostaglandins  

$$\downarrow R_A + \downarrow R_E \longrightarrow fGFR + fRBF$$

Blockade of prostaglandin synthesis  $\rightarrow \downarrow$  GFR

- This is usually important only when there are other disturbances that are already tending to lower GFR
- e.g. nonsteroidal antiinflammatory drugs in a
- volume depleted patient, or a patient with heart failure,
- cirrhosis, etc

## 4. Endothelial-Derived Nitric Oxide (EDRF) $\downarrow R_A + \downarrow R_E \longrightarrow \uparrow GFR + \uparrow RBF$

- Protects against excessive vasoconstriction
- Patients with endothelial dysfunction (e.g. atherosclerosis) may have greater risk for excessive decrease in GFR in response to stimuli such as volume depletion

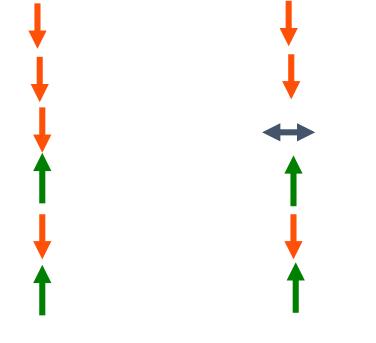
- Hepatorenal syndrome decreased renal function in cirrhosis or liver disease?
- Acute renal failure (e.g. contrast media nephropathy)?
- Hypertensive patients with chronic renal failure?

Endothelin antagonists may be useful in these conditions

# Summary of neurohumoral control of GFR and renal blood flow

### Effect on RBF Effect on GFR

Sympathetic activity Catecholamines Angiotensin II EDRF (NO) Endothelin Prostaglandins



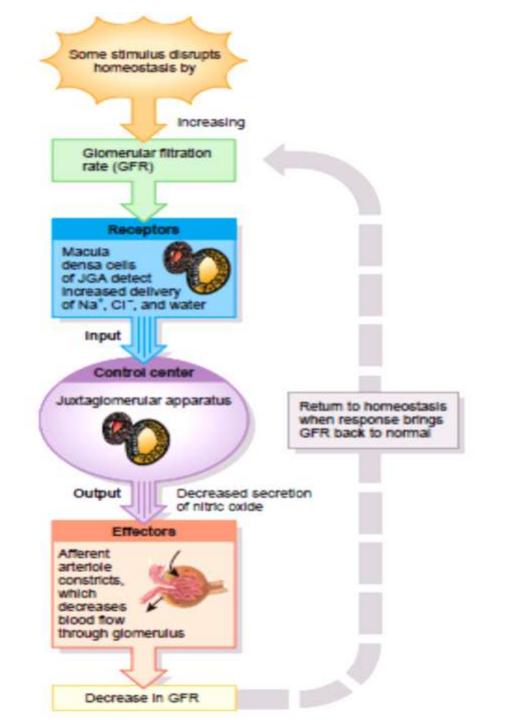
↑ increase ↓ decrease → no change

- 7. Autoregulation of GFR and Renal Blood Flow
  - Myogenic Mechanism
  - Macula Densa Feedback
    - (tubuloglomerular feedback)
  - Angiotensin II ( contributes to GFR but not RBF autoregulation)

# Renal Autoregulation of GFR

# 2. Tubuloglomerular feed back mechanism:

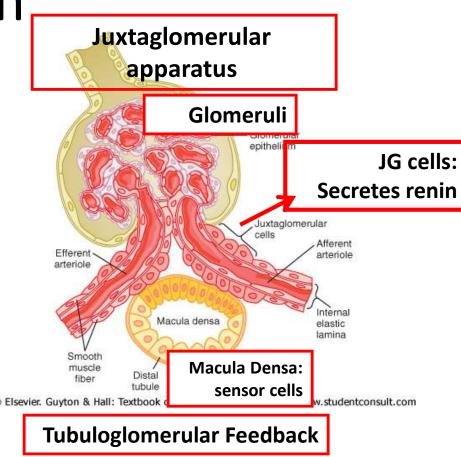
- Feedback loop consists of a flow rate (increased NaCl in filtrate) sensing mechanism in macula densa of juxtaglomerular apparatus (JGA)
- Increased GFR (& RBF) inhibits release of the vasodilator ; Nitric Oxide (NO)



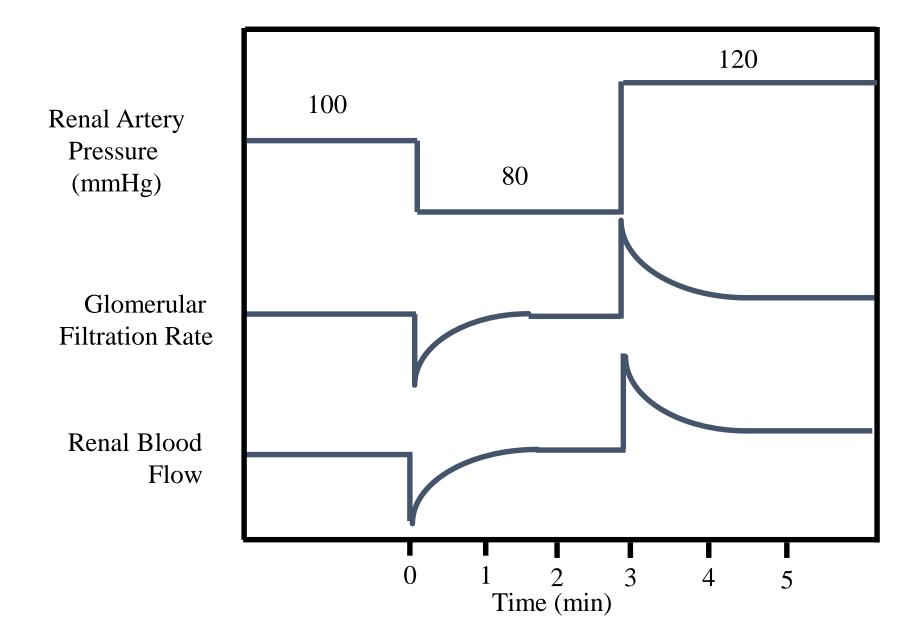
# **Renin secretion regulation**

#### **1- Perfusion Pressure**

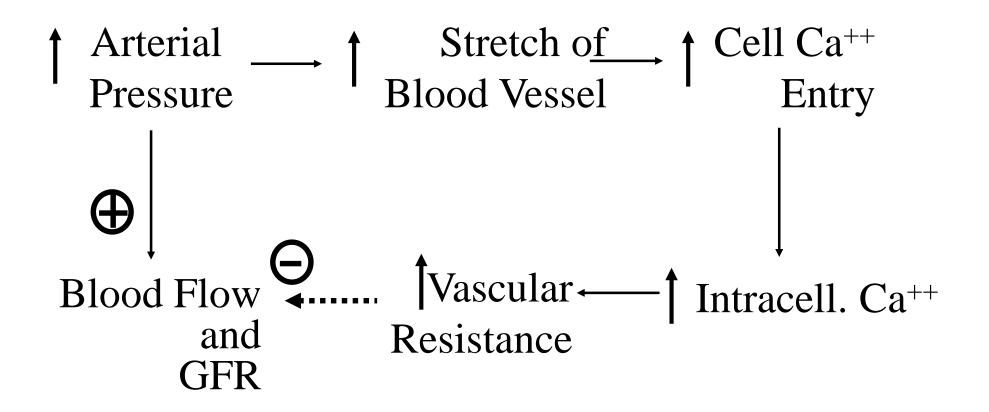
- low perfusion in afferent arterioles stimulates renin secretion while hig perfusion inhibits renin secretion.
- 2-Sympathetic nerve activity
- Activation of the sympathetic nerve fibers in the afferent arterioles increases renin secretion.
- 3- NaCl delivery to macula densa: When NaCl is decreased, Renin secretion is stimulated and vice versa. (Tubuloglomerular Feedback)



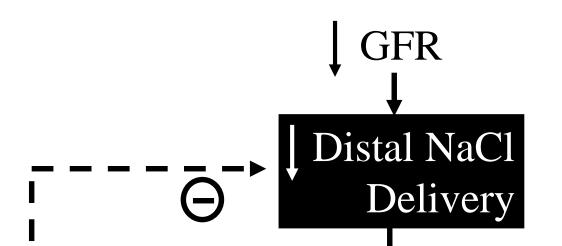
## **Renal Autoregulation**



## **Myogenic Mechanism**

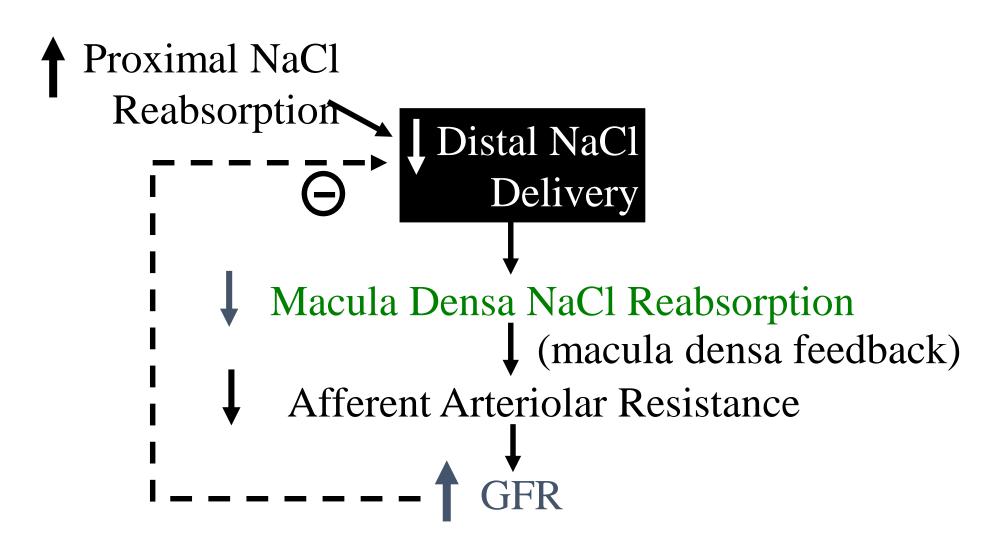


## Macula Densa Feedback

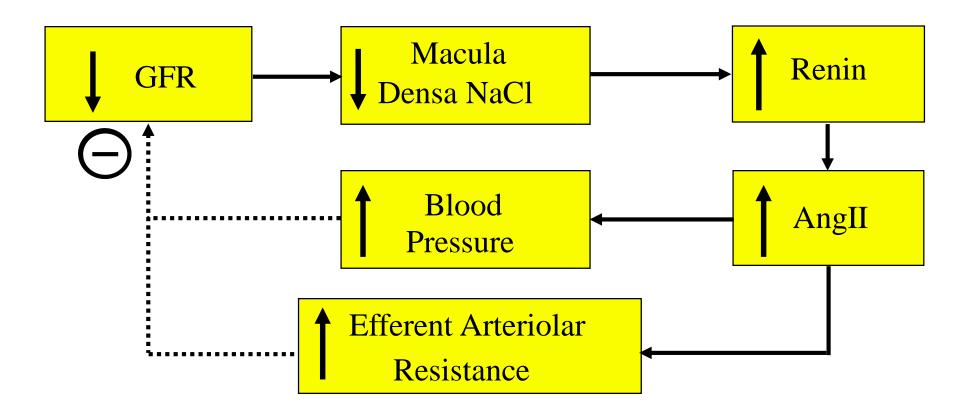


Macula Densa NaCl Reabsorption
 (macula densa feedback)
 Afferent Arteriolar Resistance

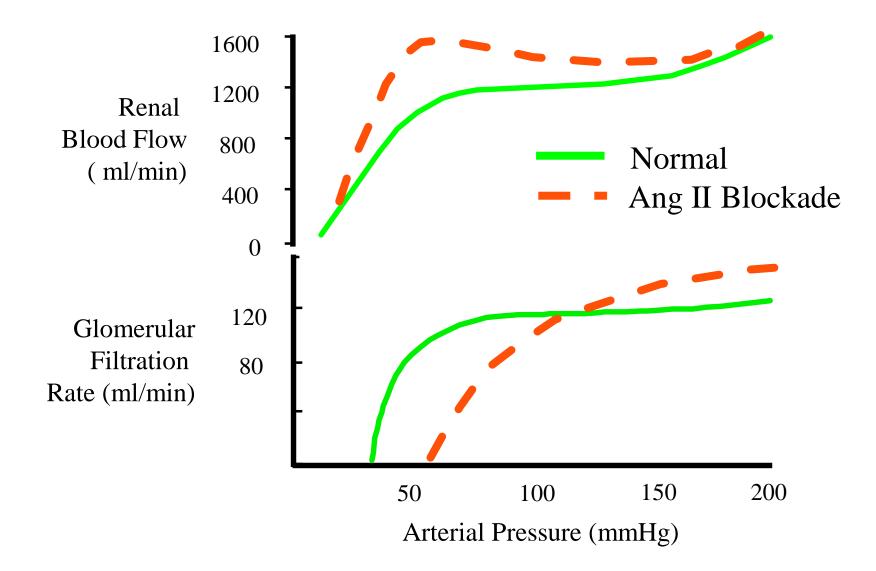
## Macula Densa Feedback

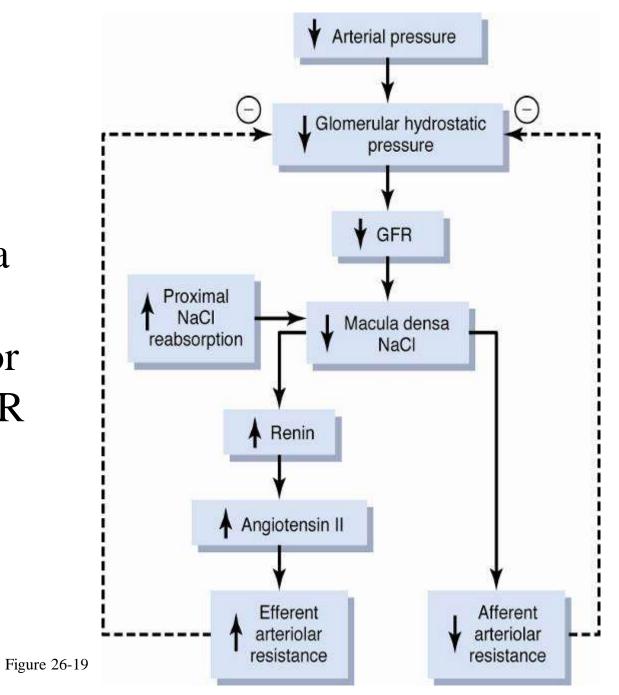


## **Regulation of GFR by Ang II**



### Ang II Blockade Impairs GFR Autoregulation

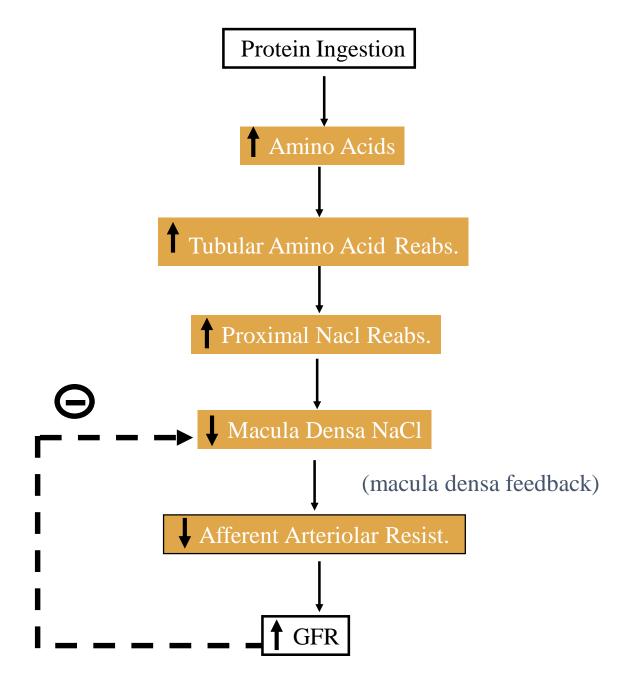




Macula densa feedback mechanism for regulating GFR

### **Other Factors That Influence GFR**

- Fever, pyrogens: increase GFR
- Glucorticoids: increase GFR
- Aging: decreases GFR 10% / decade after 40 yrs
- Hyperglycemia: increases GFR (diabetes mellitus)
- Dietary protein: high protein increases GFR low protein decreases GFR



## **Importance of Autoregulation**

Arterial Pressure	GFR	Reabsorption	Urine Volume
Poor Autoregulation + no change in tubular reabsorption			
100	125	124	1.0
120	150	124	26.0 = 37.4  L/day!
Good Autoregulation + no change in tubular reabsorption			
120	130	124	5.0
Good Autoregulation+adaptive increase in tubular reabsorption			
120	130	128.8	1.2