



The purpose of education is to replace an empty mind with an open one.

#### Malcolm S. Forbes

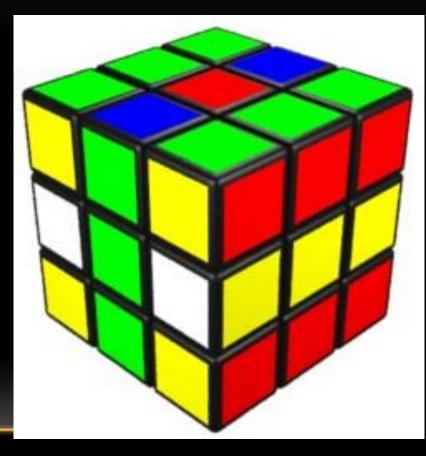
لا يسمح بالخول بعد بدء المحاضرة أو تغيير الشعبة لأي سبب كان.

لايسمح بتسجيل المحاضرات

سيحرم من الامتحان النهائي كل من يتجاوز غيابه 15% من المحاضرات

## Remember !!!!

Histology is a 2 dimensional study of a 3 dimensional reality.





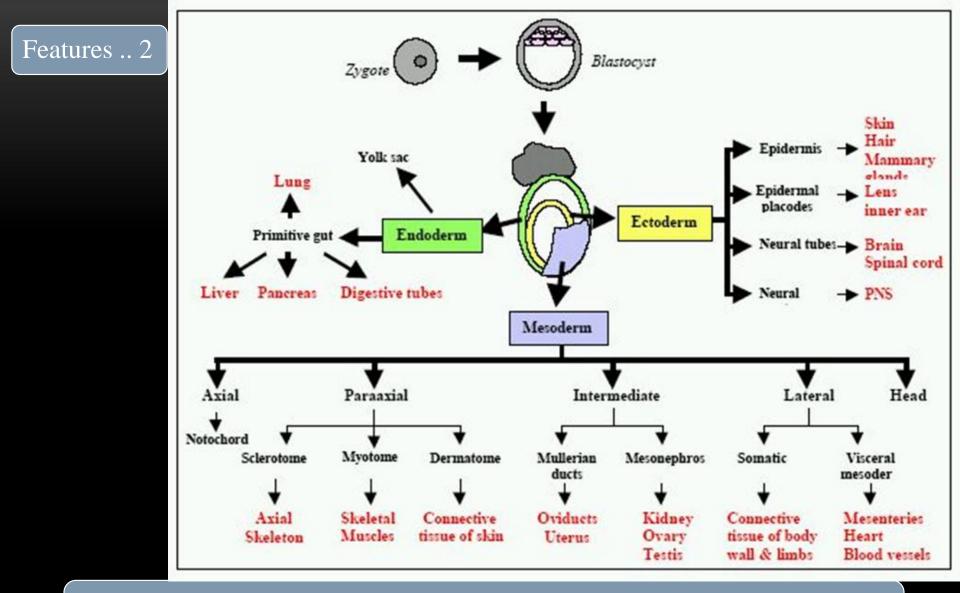


# It is the interface between all tissues and the external and internal environment of the body.

# Features

Always on top: covers a surface OR lines a cavity



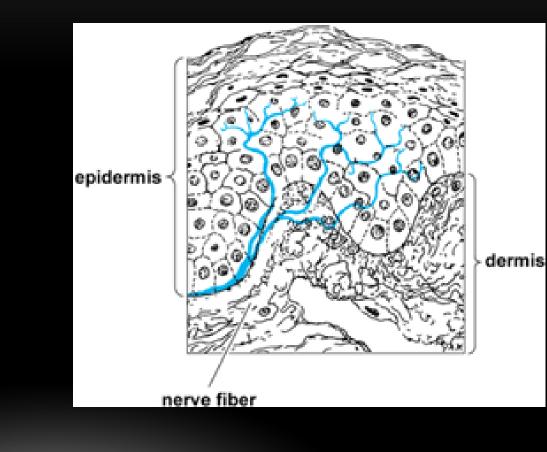


Originates from any of the three embryonic layers.

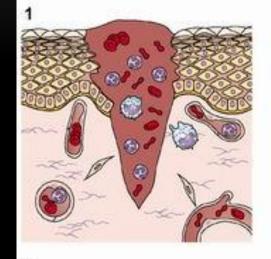


Avascular and depends in its nutrition on perfusion from the underlying connective tissue.

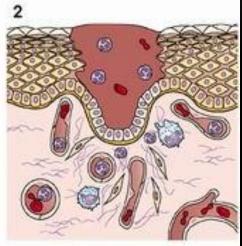
Richly innervated. Free nerve endings can be seen between epithelial cells.



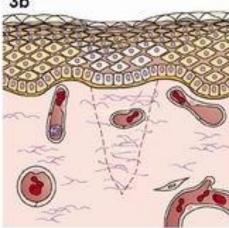
### Has a high regenerative power.



3a



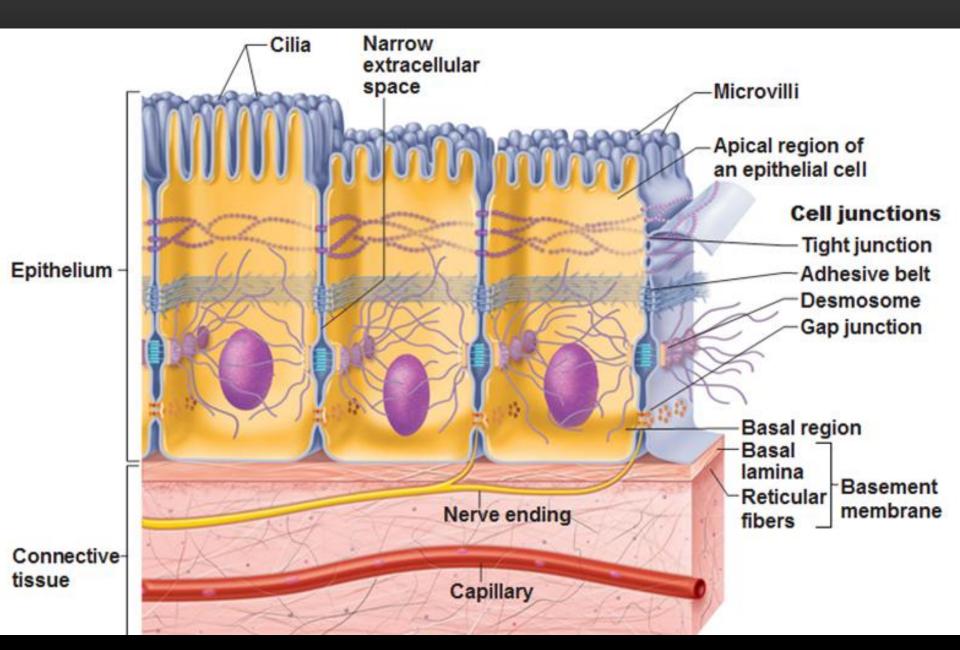
3b



Cells are closely packed forming sheets or membranes.

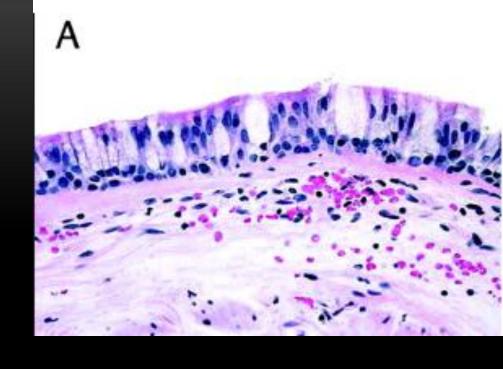


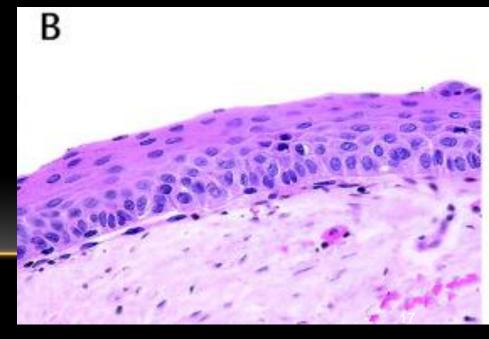
Cells show polarity; organelles are collected at the site of function.



## Feature 8

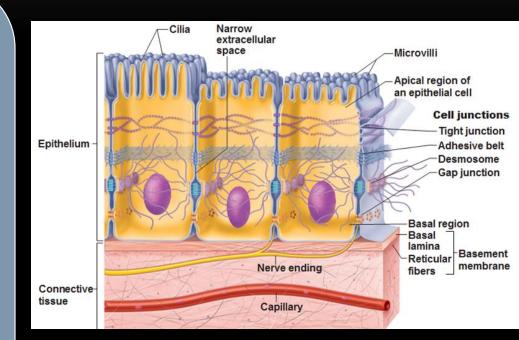
Metaplasia = reversible change of one type of epithelium to another in response to a stimulus





## Feature 9

Always rests on a "Basement Membrane or Basal Lamina"



# Lecture 2

You do not have to burn books to destroy a culture. Just get people to stop reading them. Ray Bradbury

# Basal Laminae & Basement Membrane

### Basal Lamina

### Only visible with E.M

### Found also in other tissues

Components are secreted by epithelium, connective tissue, muscle, Schwann cell

## Layers of Basal Lamina

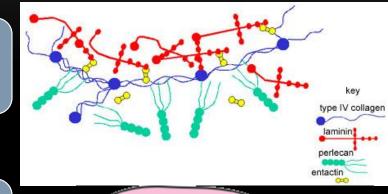
#### Layers of the Basal Lamina

- Lamina Lucida
- Lamina Densa

## Lamina Reticularis: not part of the basal lamina

## Molecular components are variable but include:

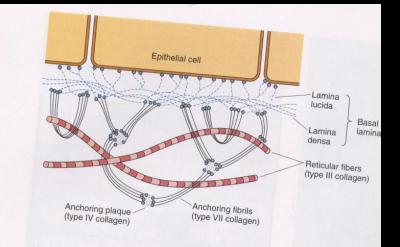
- type IV collagen,
- Glycoproteins (Laminin, entactin...)
- Proteoglycans (Perlecan)











## Functions of Basal lamina

Support

Selective barrier

Influencing cell polarity

Regulation of proliferation and growth

Affect cellular metabolism

Affect cell-cell interaction

### Clinical Importance of Basal Lamina

# Tissue culture

# Tumor grading

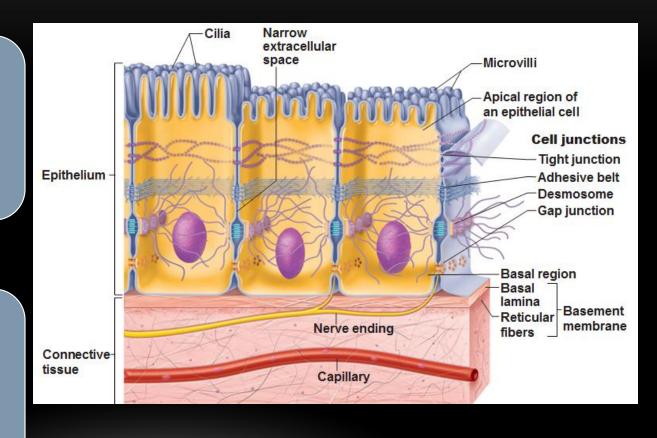
26

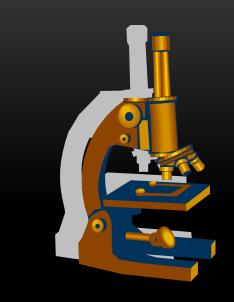
### The epithelial cell

#### Shown to have the following domains:

- Apical
- Baso-lateral

Each domain shows modifications to suit its functions.





## Fact.... Most of the tumors after the age of 45 are of epithelial origin.

# **Functions of Epithelium**

Protection:

Transcellular transport:

Secretion:

Absorption:

Selective permeability:

Detection of sensations:

### **Classification of Epithelium**

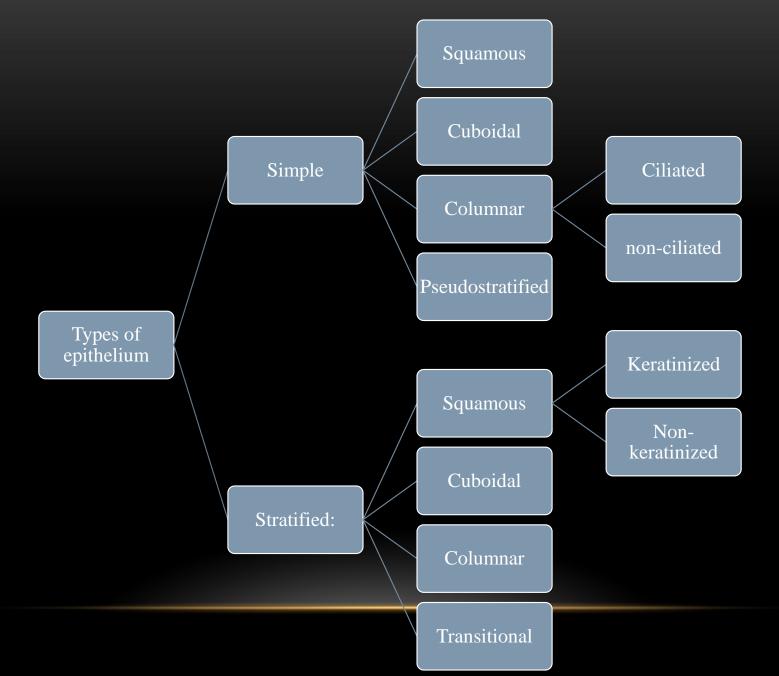
According to the number of cell layers above the basal lamina epithelium is classified into:

- simple, or
- stratified

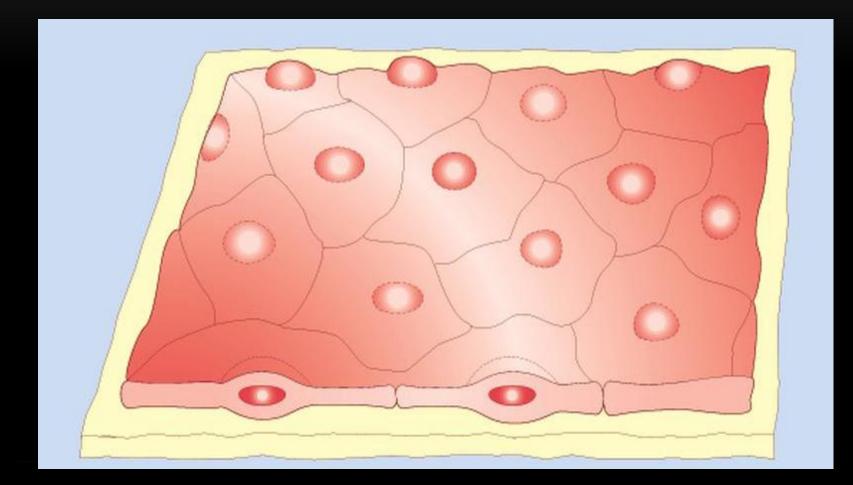
Simple epithelium is named according to the shape of its cells.

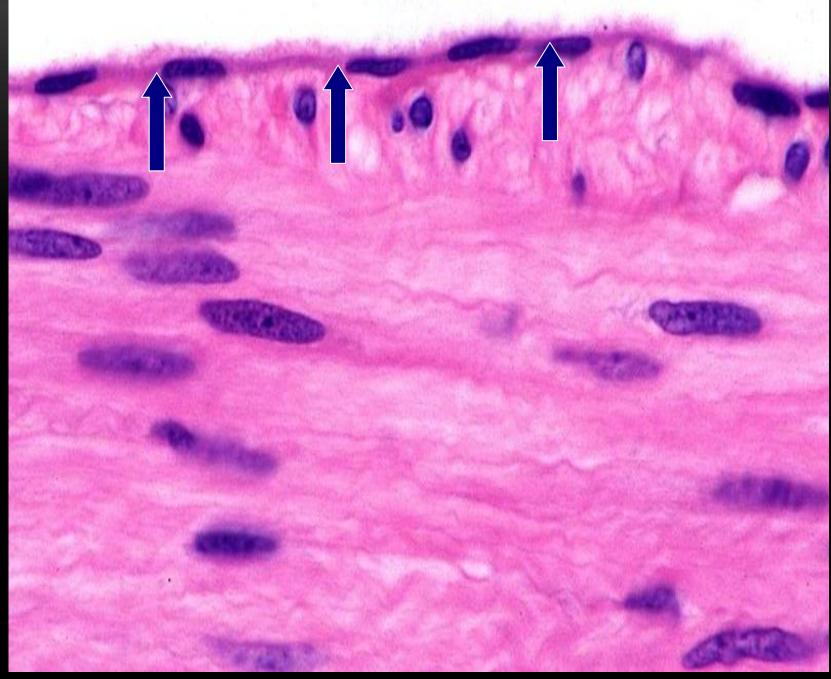
Stratified epithelium is named according to the shape of the cells in the outermost layer.

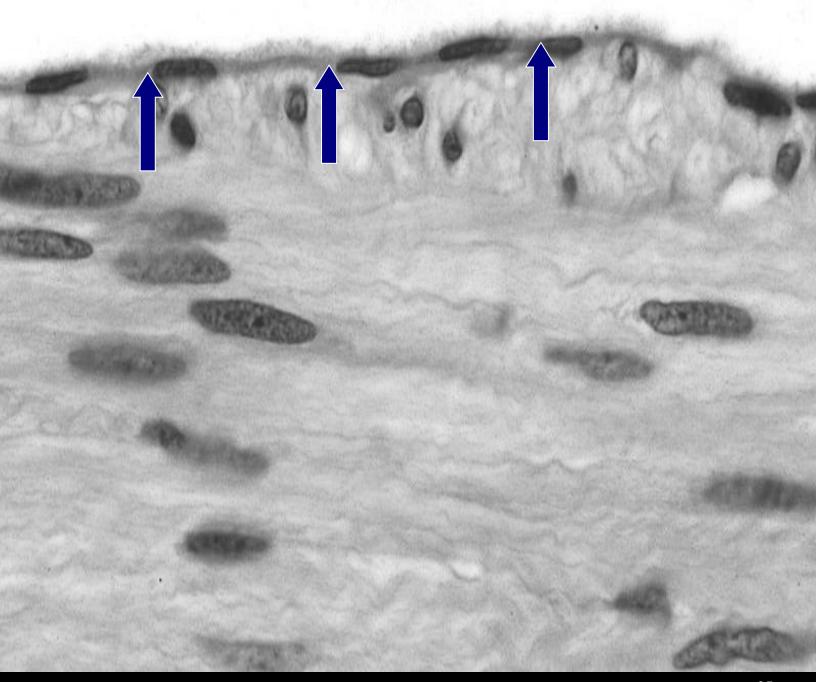
## Today a reader, tomorrow a leader. Margaret Fuller



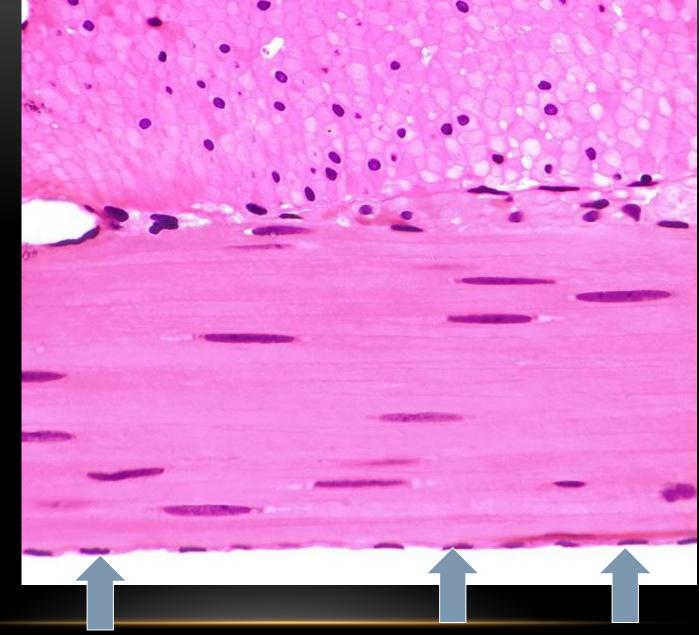
### **Simple Squamous Epithelium**



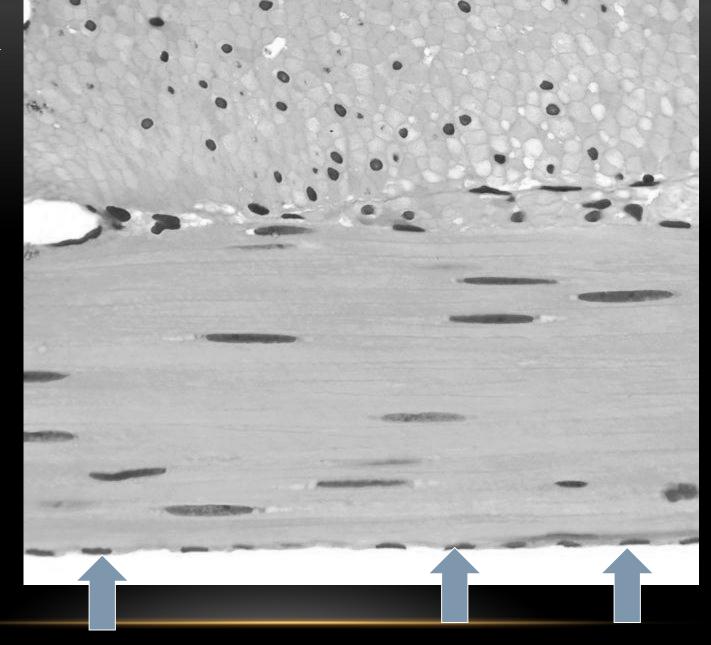




#### Mesothelium



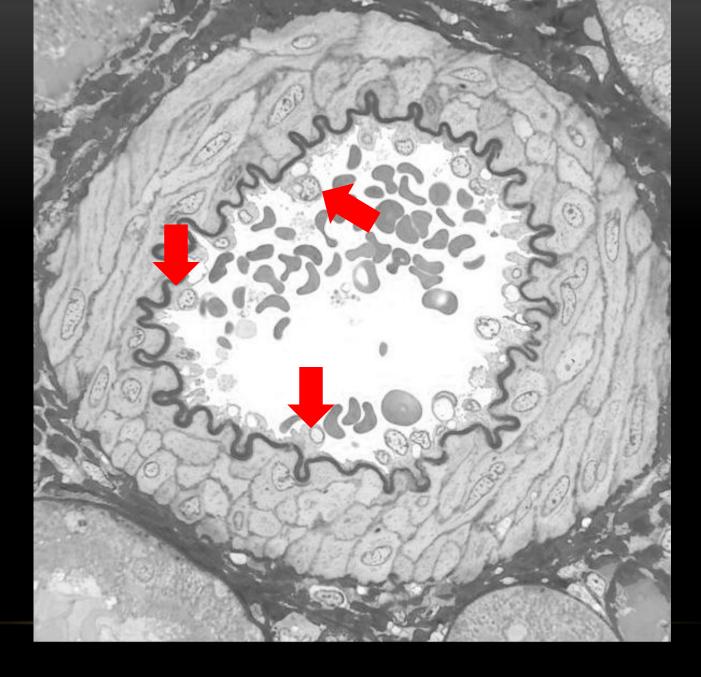
#### Mesothelium



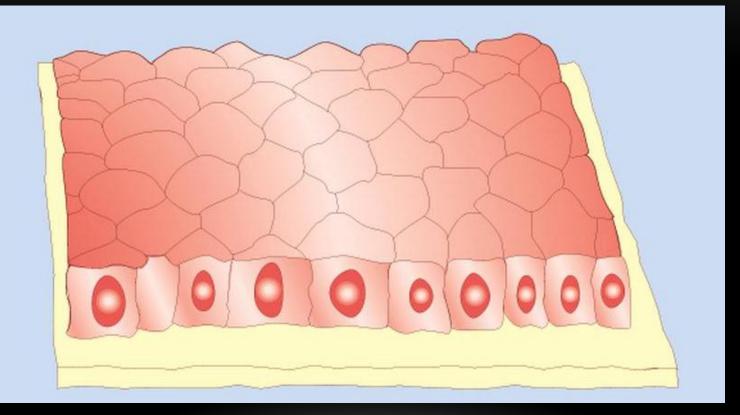
#### Endothelium



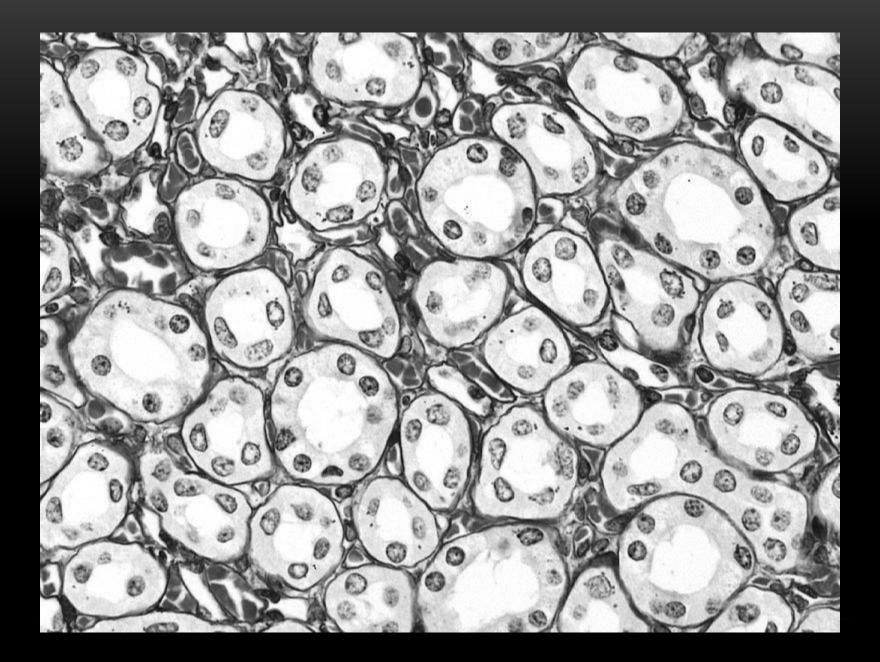
#### Endothelium

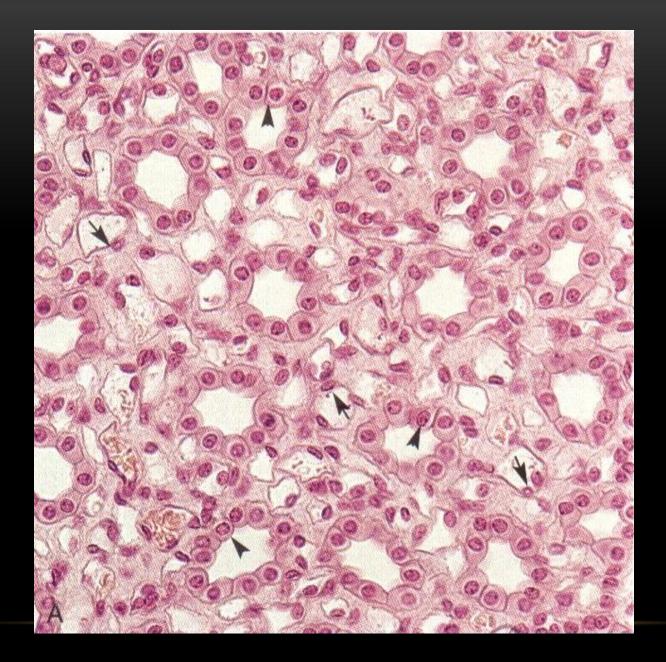


# **Simple Cuboidal**

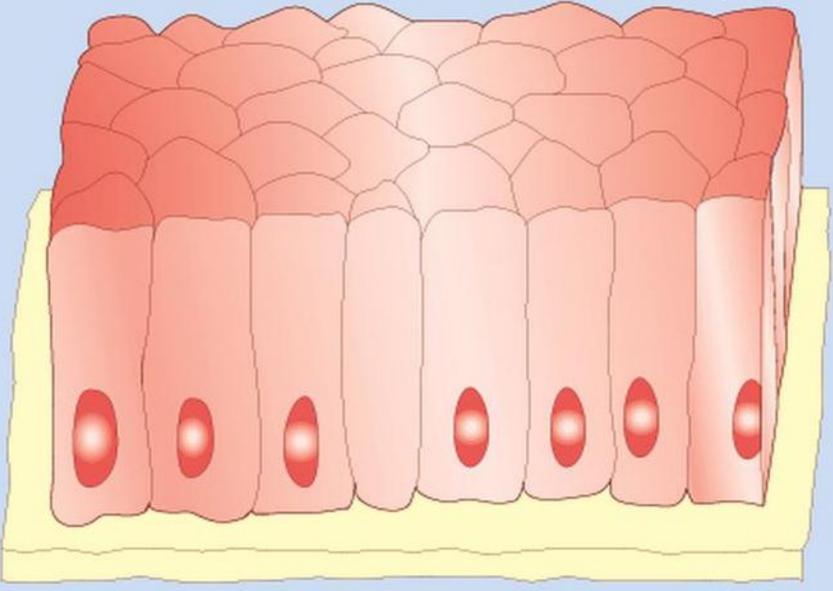


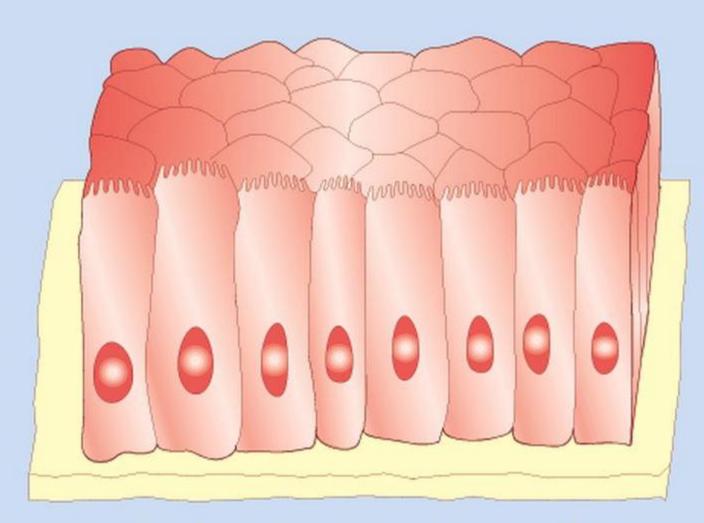




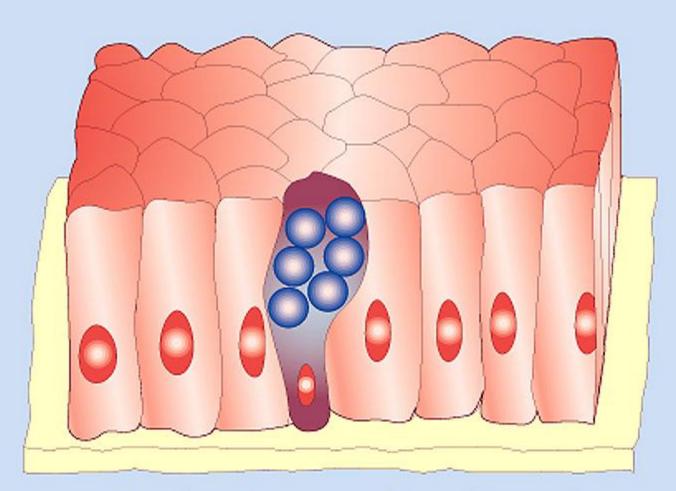


#### Simple columnar

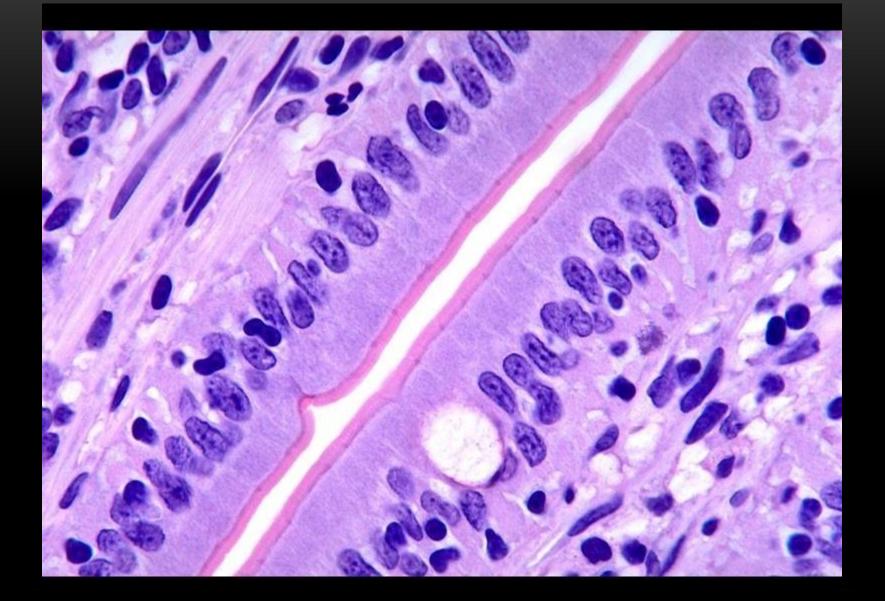


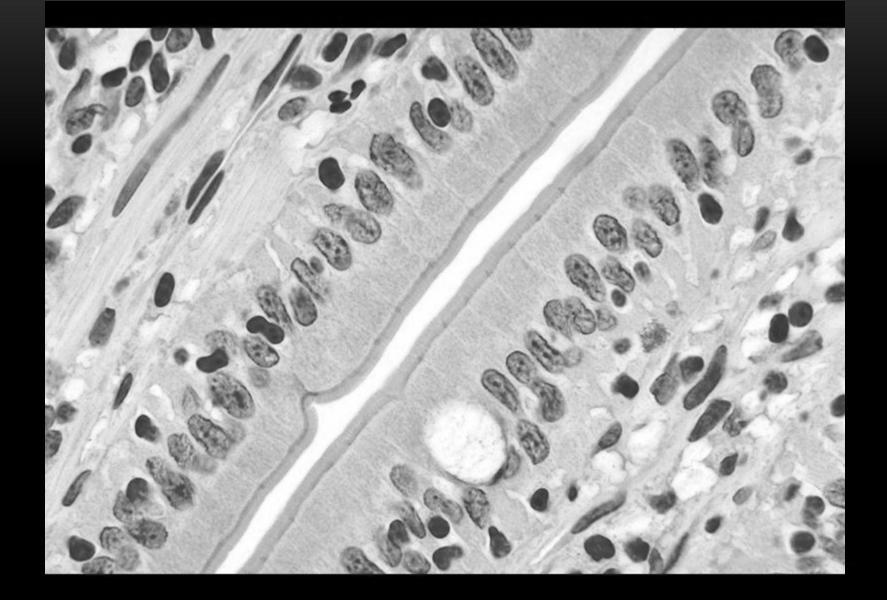


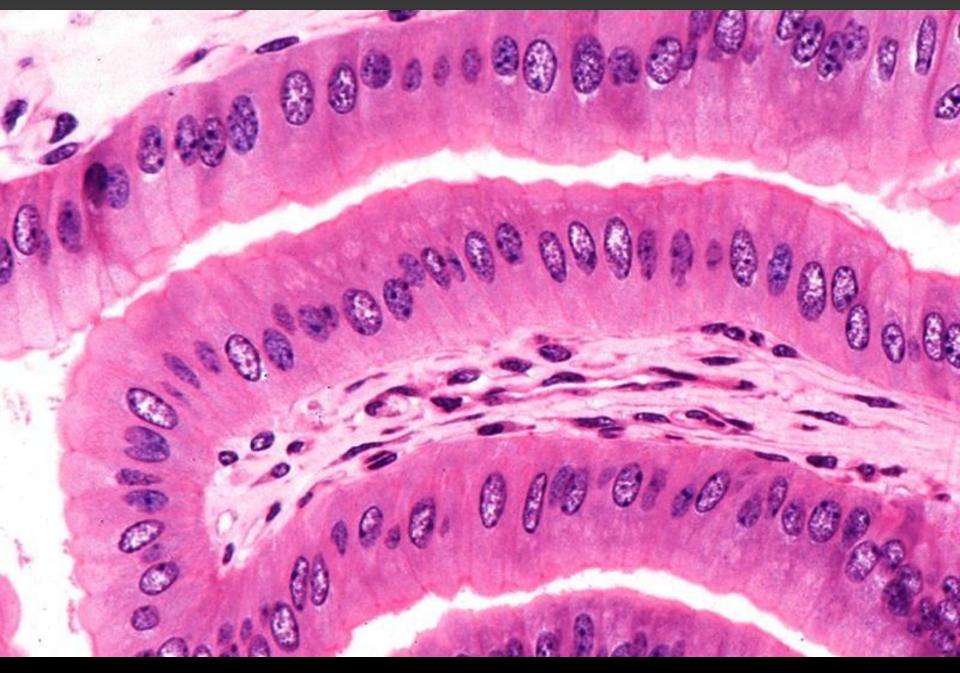
### Simple columnar with microvilli

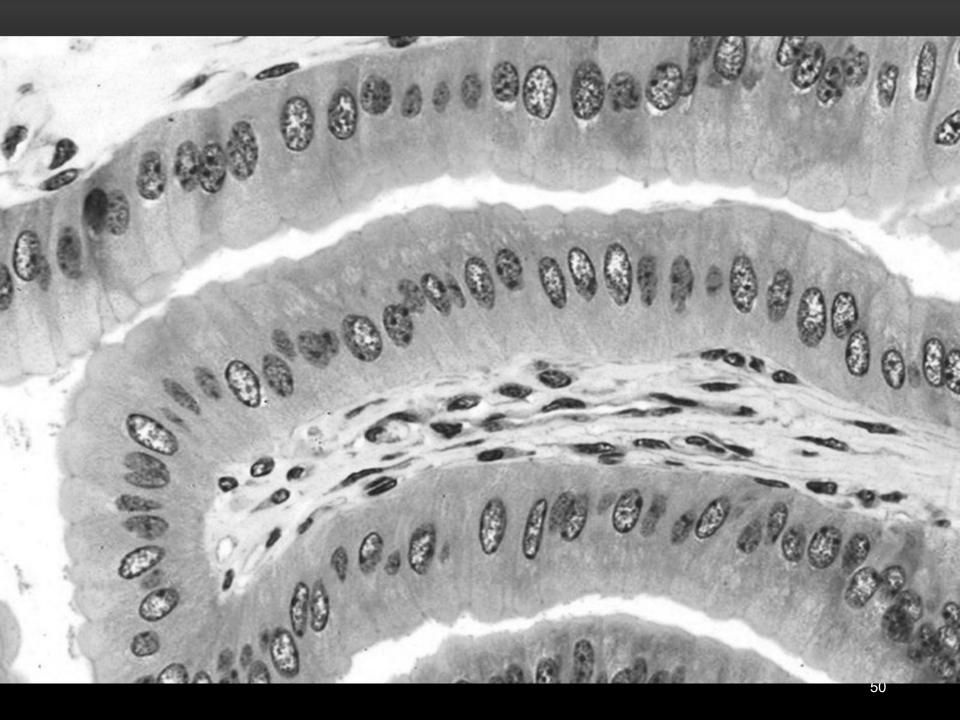


## Simple columnar with goblet cells

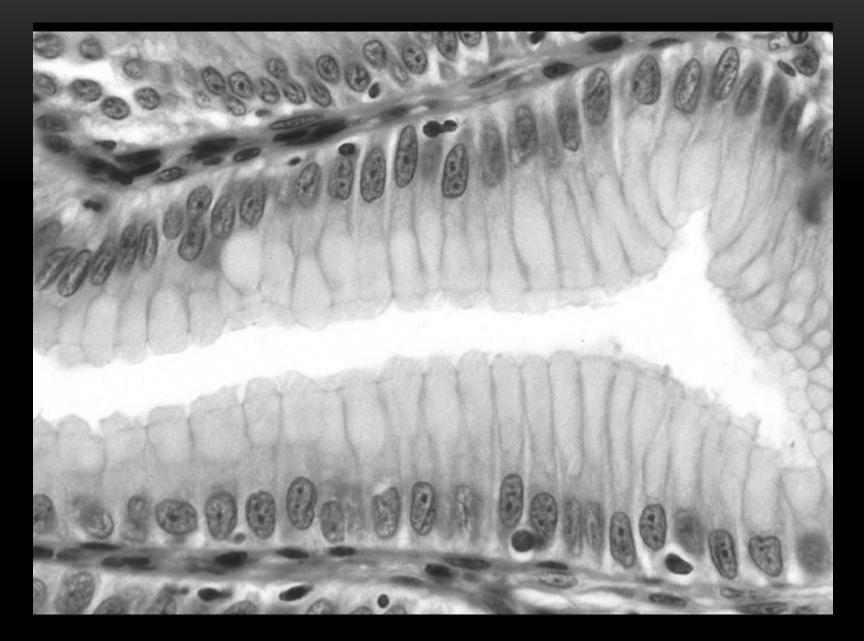


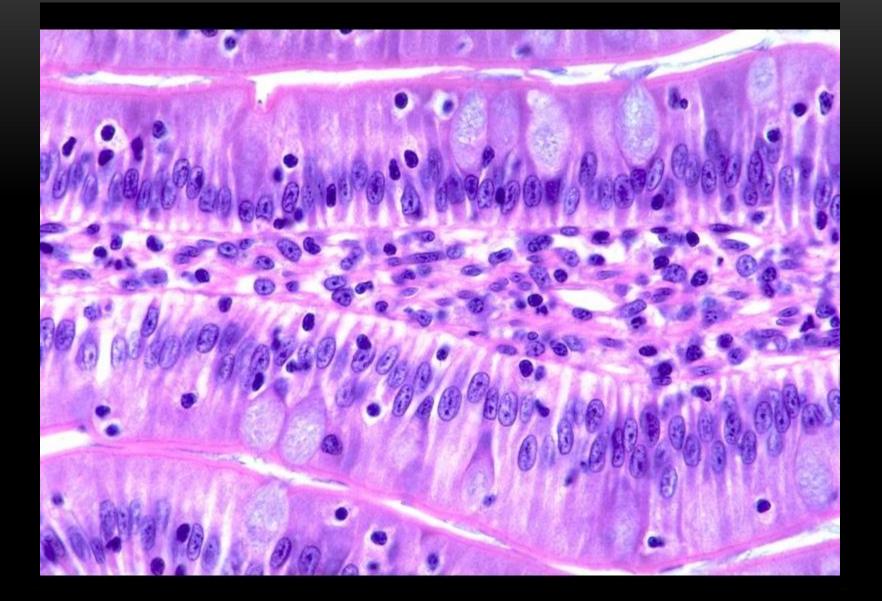


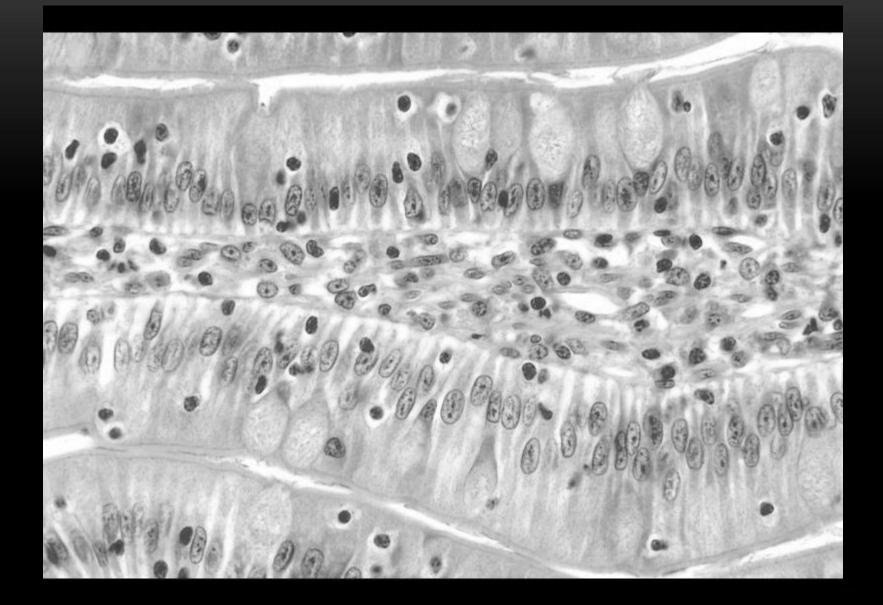


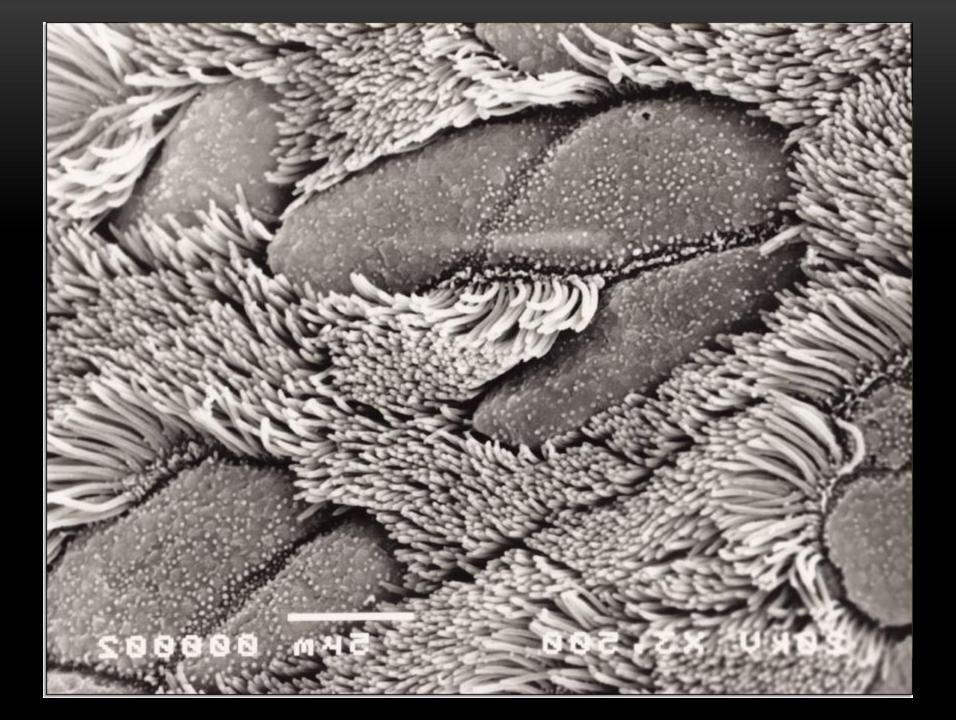


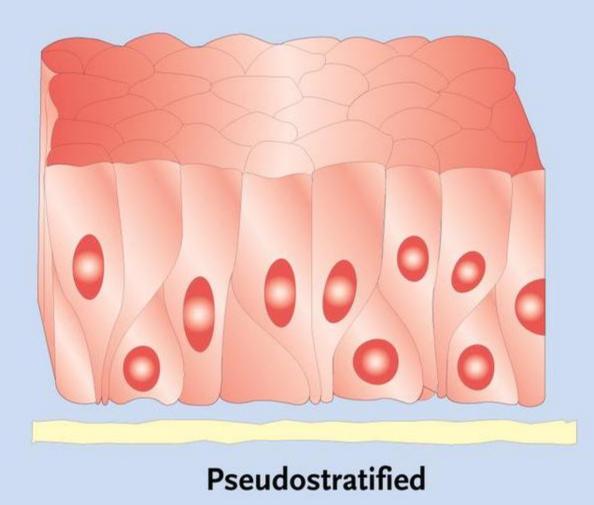








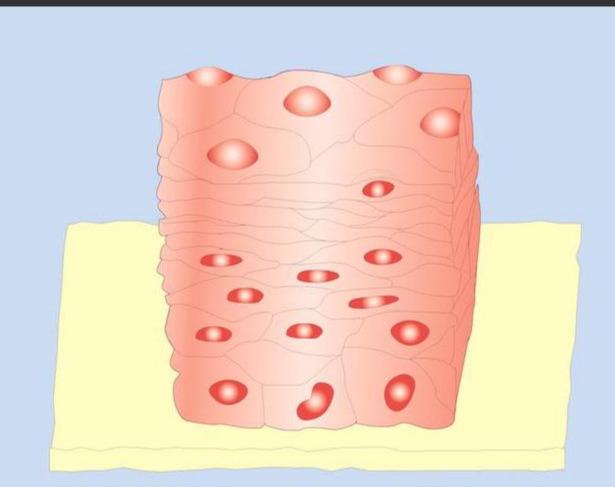






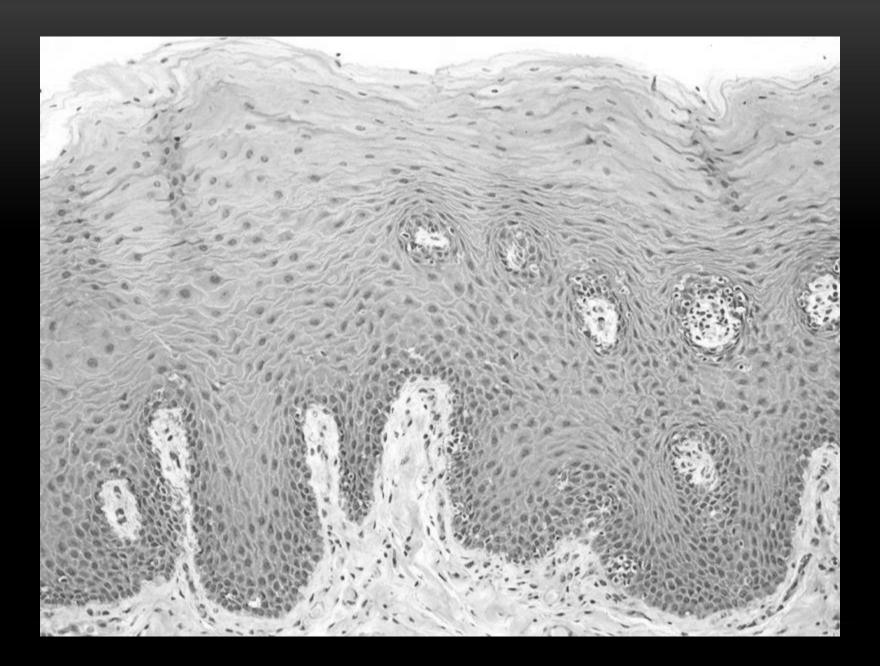
People call it luck when you've acted more sensibly than they have.

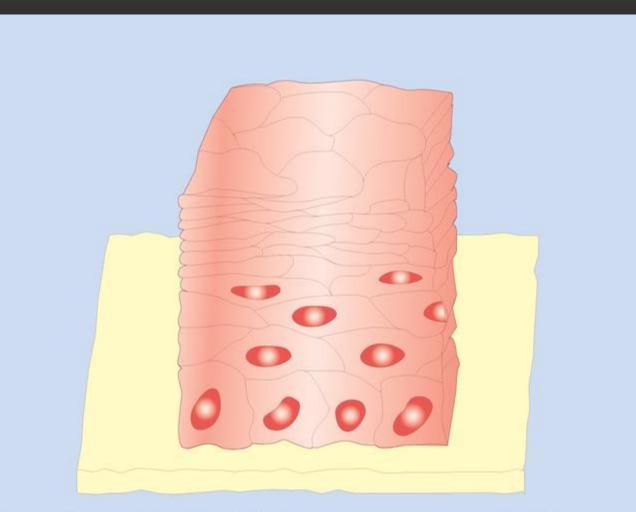
Ann Tyler



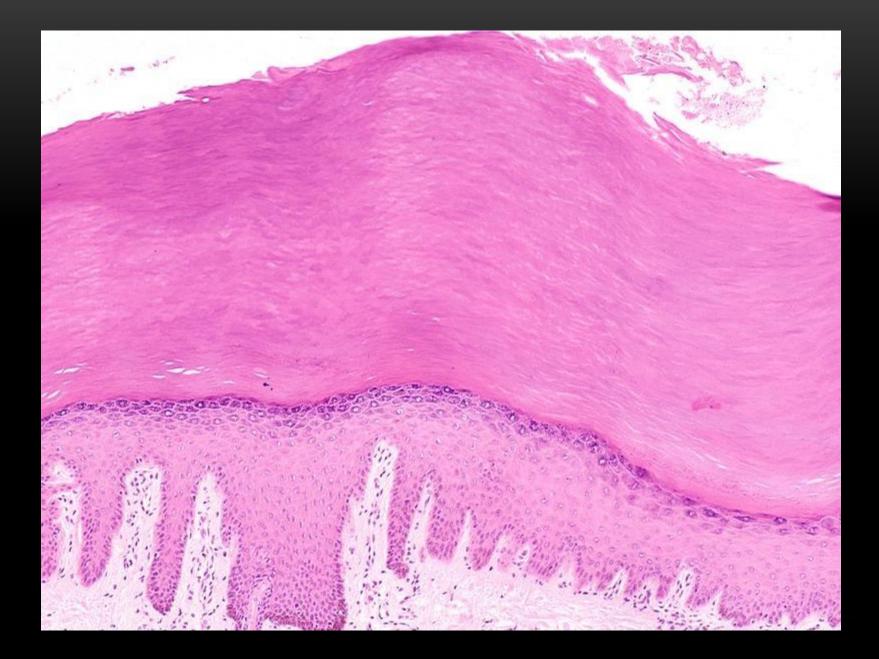
### Stratified squamous epithelium

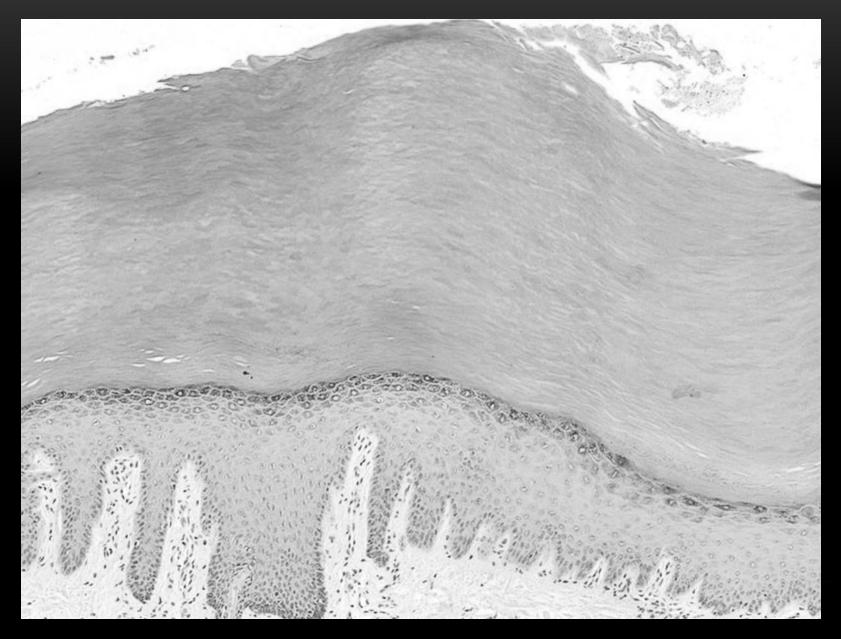




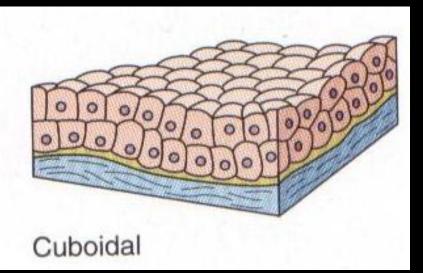


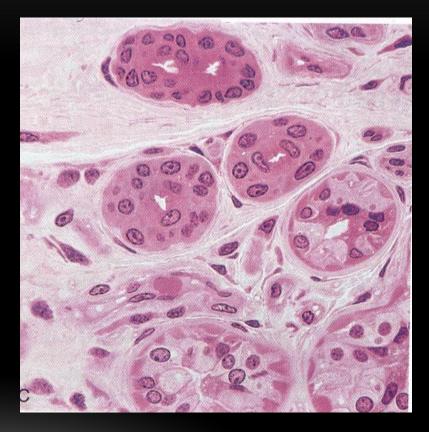
### Keratinized stratified squamous epithelium



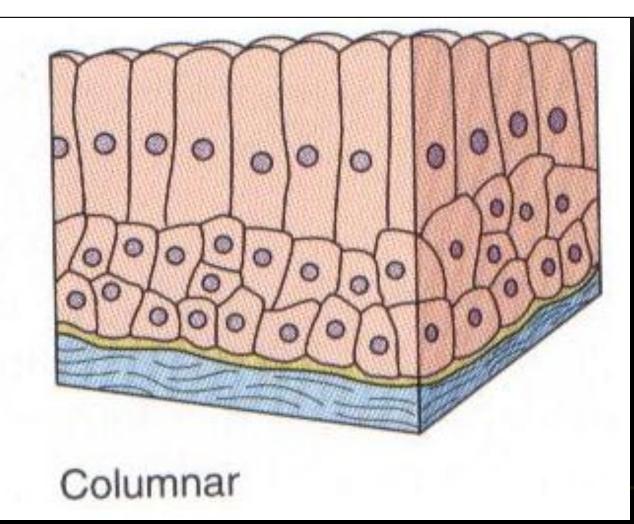


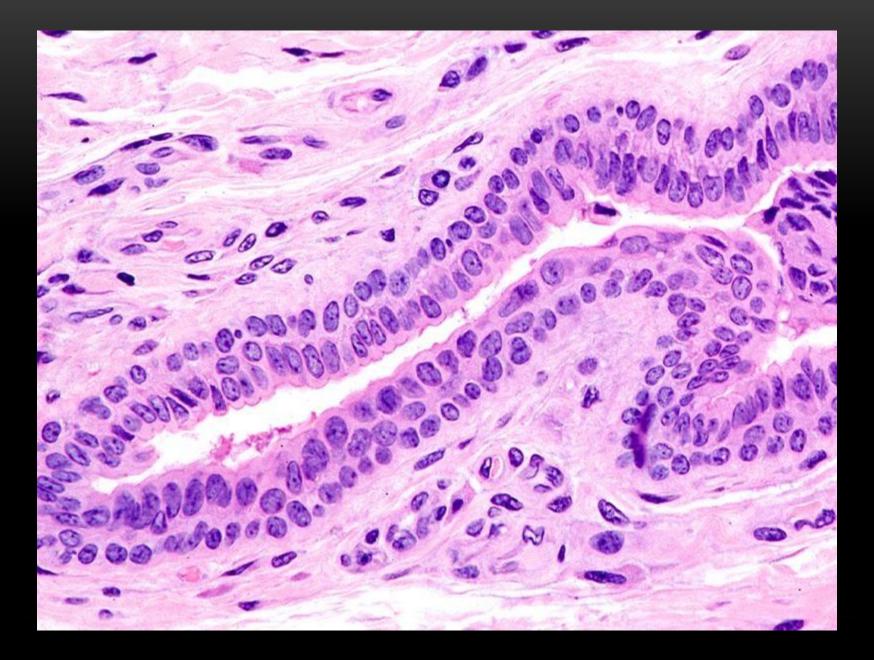
# **Stratified Cuboidal Epithelium**



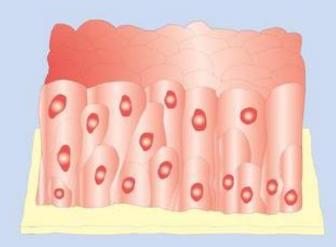


# **Stratified Columnar Epithelium**

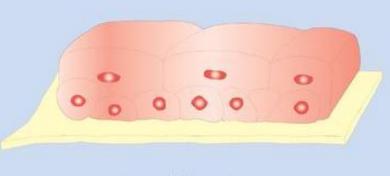




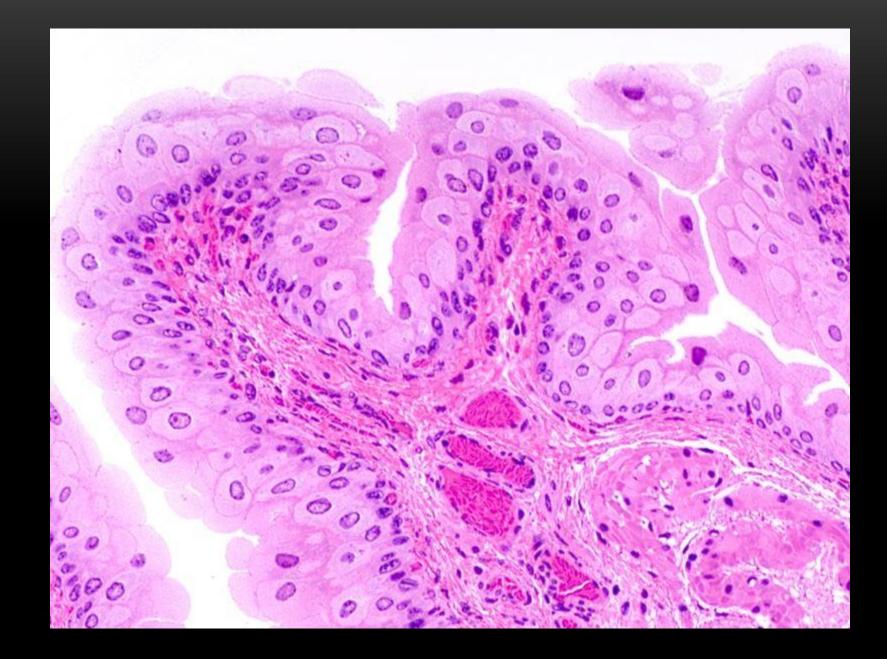




Transitional relaxed



Transitional stretched









The saddest aspect of life right now is that science gathers knowledge faster than society gathers wisdom.

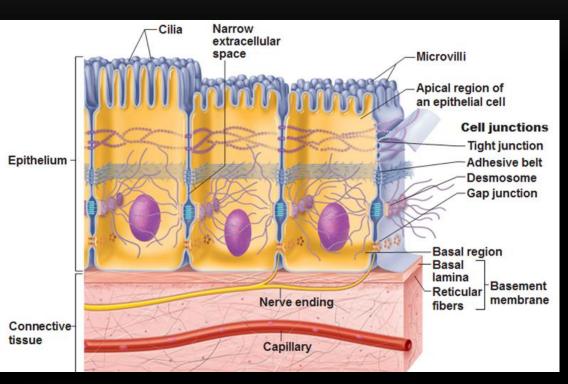
Isaac Asimov

## Apical Domain

It is the part of the cell that faces the lumen (the free surface of the cell).

It is rich in ion channels, carrier proteins and hydrolytic enzymes.

The apical modifications are:



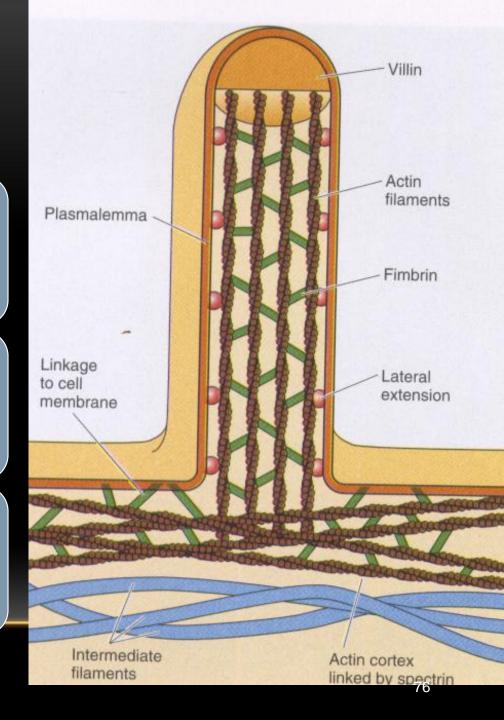
- Microvilli.
- Stereocilia.
- Cilia.
- Flagella.

## Microvilli

Present mainly in absorptive cells.

Their number and size vary according to the degree of activity of the cell.

They are usually crowded on the cell apex forming the <u>striate</u> <u>border</u> in the intestine and the <u>brush border</u> in the kidney.



### **Structure of the Microvillus**

The microvillus is 1-  $2\mu$  in length.

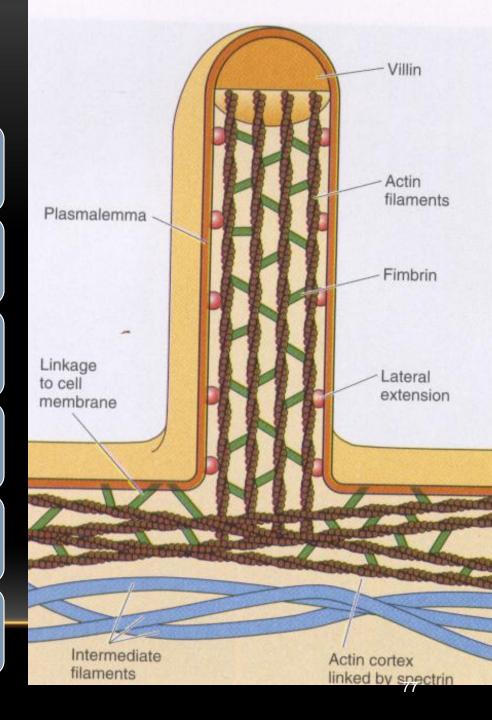
Contains a core of 25-30 actin filaments.

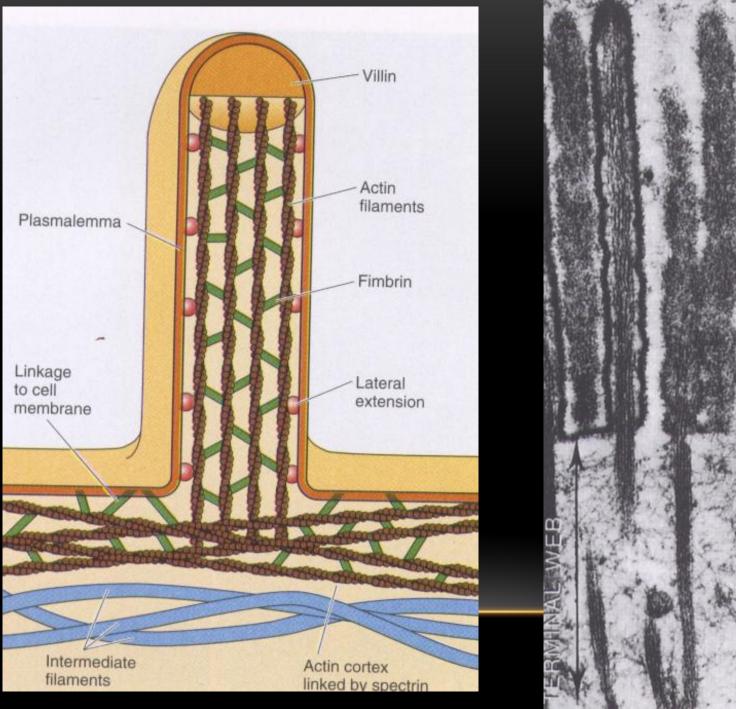
Actin filaments are cross-linked with villin.

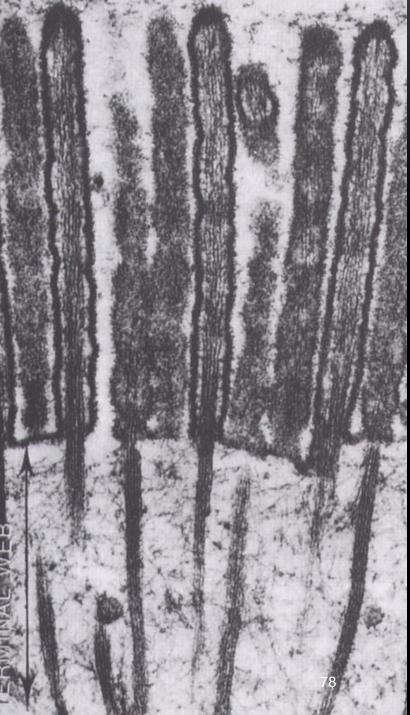
The actin filaments are inserted into the *terminal web*.

The terminal web is a network of actin and spectrin supported by myosin, IF, and camodulin in the apical part of the cell.

The microvillus is covered by <u>*glcocalyx*</u>; it gives PAS +ve reaction

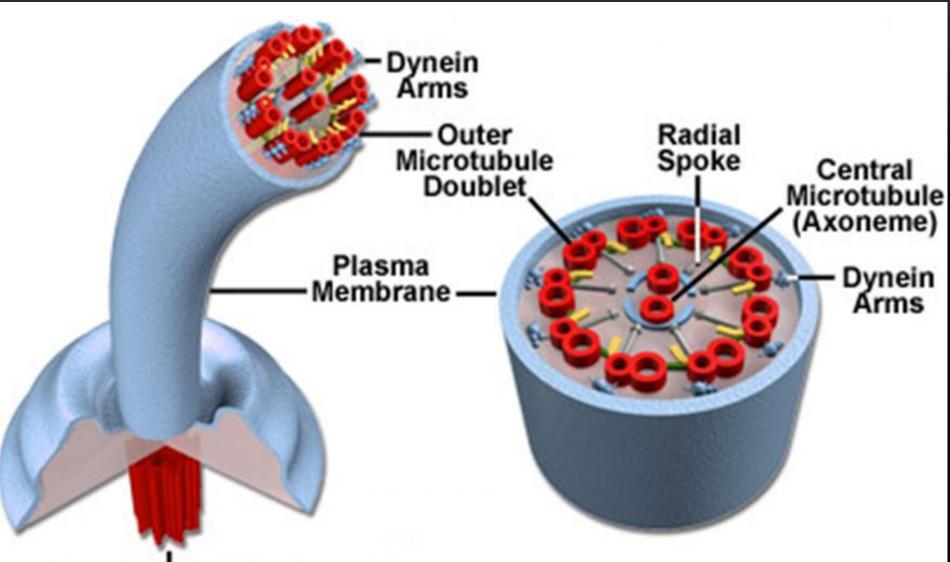








Stereocilia are long immotile microvilli present in the epididymis and inner ear. They have special functions in these places.



Basal Body (Kinetosome)

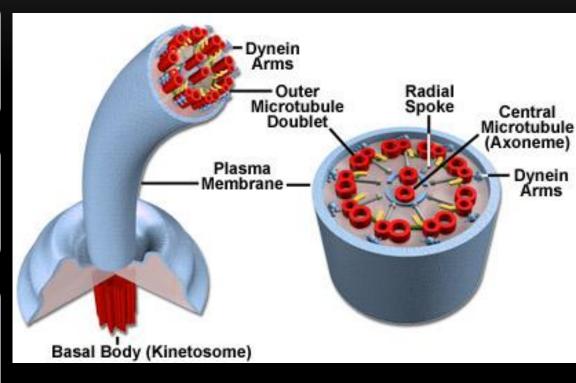
## Cilia

Motile cytoplasmic hair like projections capable of moving fluid and particles along epithelial surfaces.

<u>Measurements</u>: length 7-10 $\mu$ , diameter 0.2  $\mu$ .

Number of cilia/cell is variable and ranges 1-300 cilium/cell.

They move rhythmically and rapidly in one direction.



### The core of the cilium is called *axoneme*.

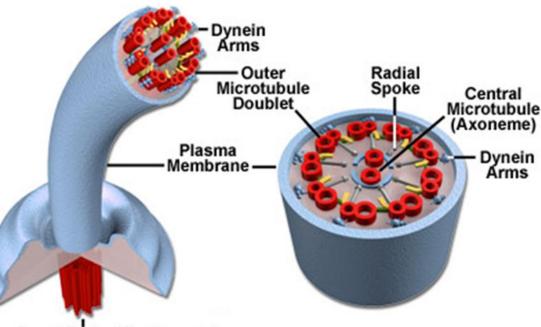
The axoneme consists of longitudinal microtubules arranged as 9 (doublets) peripheral surrounding 2 (singlets) central (9+2).

The singlets are separated by 13 protofilaments.

The doublets are composed of 2 subunits A & B.

Subunit A is formed of 13 protofilaments.

Subunit B is formed of 10 protofilaments.



Basal Body (Kinetosome)

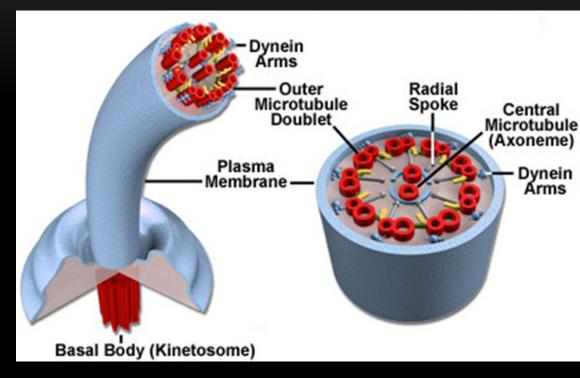
Neighboring doublets are connected by nexin.

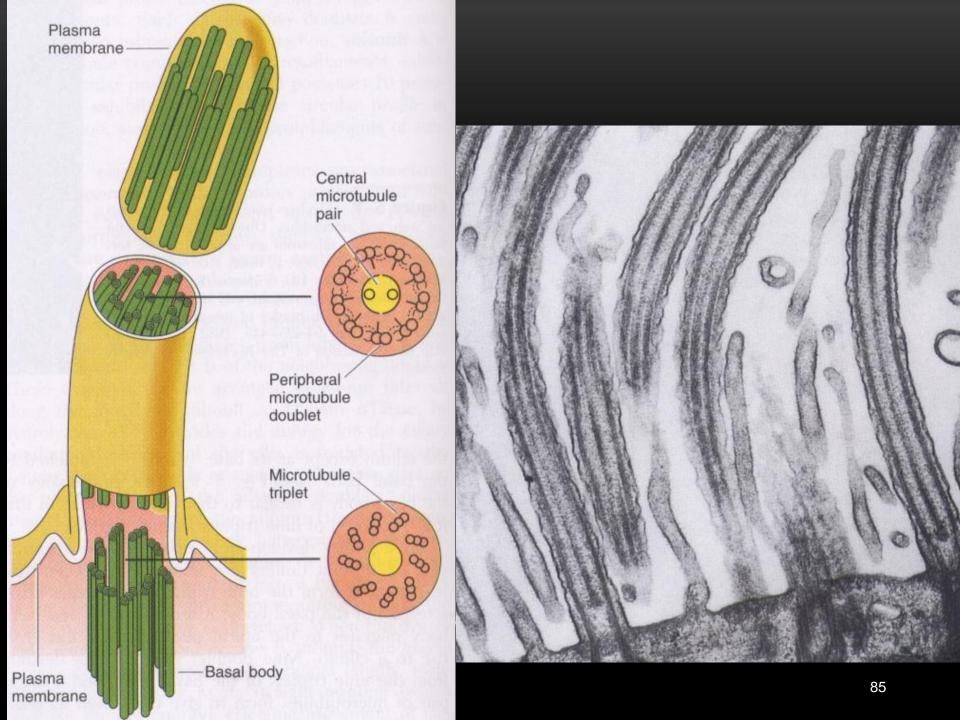
Doublets are connected to the singlets by *radial spokes*.

Dynein radiates form subunit A to subunit B.

Dynein has ATPase activity.

Cilia are attached to basal bodies similar in structure to centrioles.





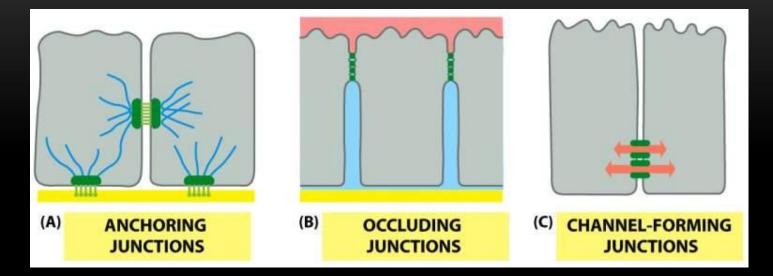
# Baso-Lateral Domain

# Terminal bars are light microscopic structures at the site of contact of cells.

# E.M revealed that the terminal bar is a junctional complex composed of:

- Occluding junctions.
- Anchoring junctions.
- Communicating junctions.

#### **Cell Junctions**

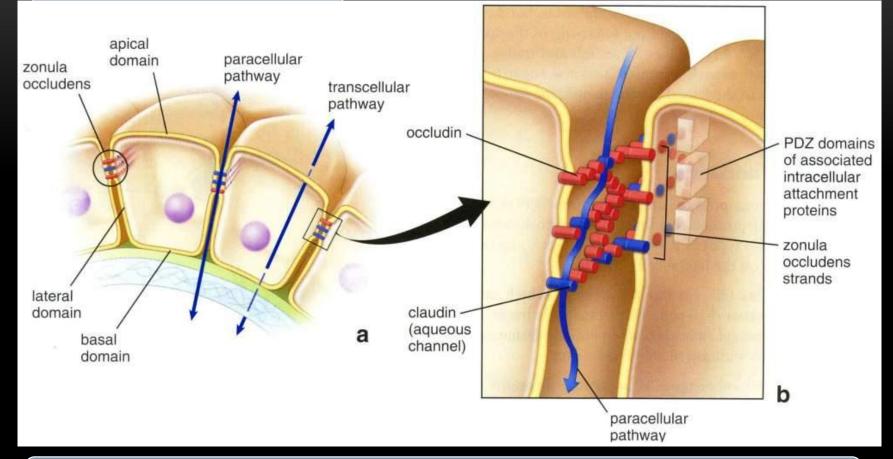


**Anchoring** (Desmosomes and Macula adherentes) - mediate cell-cell and cellmatrix adhesions; linked to cytoskeleton to transmit and distribute stress

**Occluding** (Zonula Ocludentes) - form seals between epithelial cells; block or regulate (paracellular) permeability between cells

**Channel-forming** (Gap Junction) - allow diffusion of small molecules

#### **Tight Junctions**

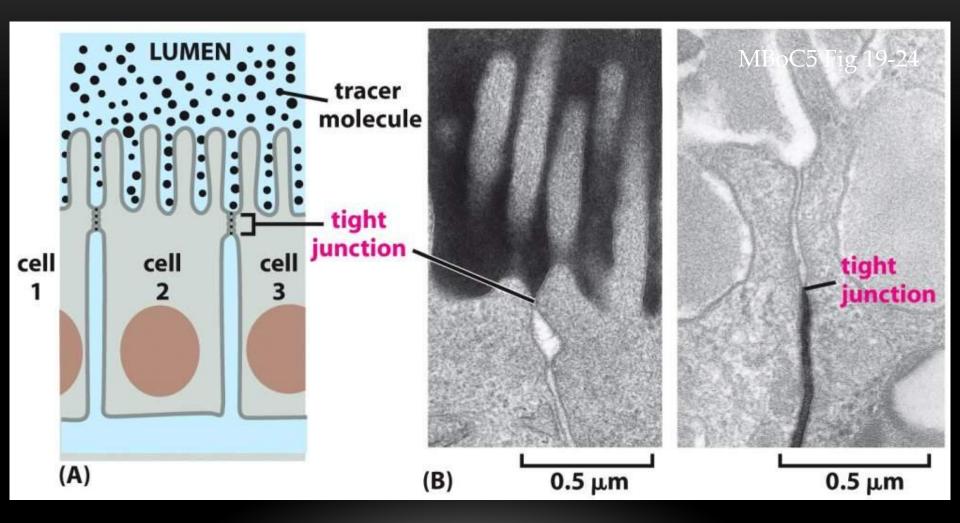


#### Occluding junction (encircles epithelial cells)

Barrier to diffusion between cells (paracellular pathway)

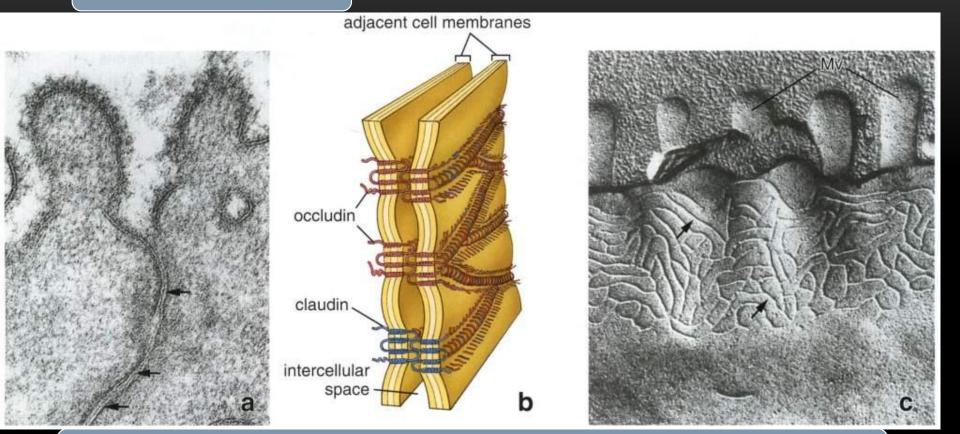
Separates apical and basolateral plasma membranes, the outer layers of 2 adjacent plasmalemma fuse together.

#### **Tight Junctions**



Tight junction blocks diffusion of soluble tracer molecules added to either the apical or basolateral compartment.

#### **Tight Junction**



#### TEM: is the most apical junction

Freeze fracture of TJ reveals ridges in membranes that correspond to sites of contact between cells

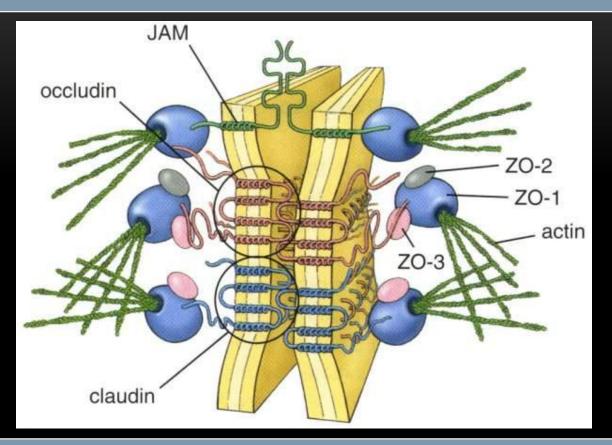
Ridges are linear arrays of occludin and claudin proteins

#### **Tight Junction Permeability**

# Top view Side view в А

Some claudins and occludins have pores (A, B, and C) that allow selective (paracellular) movement of ions or solutes

#### **Tight Junction Proteins**



Occludins and claudins are transmembrane proteins that interact across the intercellular space to form TJs

ZO (zonula occludens) proteins 1-3 link occludin and claudin to each other, to JAMs, and to actin filaments

### Tight junction

#### Adherens junction

#### Desmosome

Microvillus

#### Gap junction -

## **Zonula Adherens**

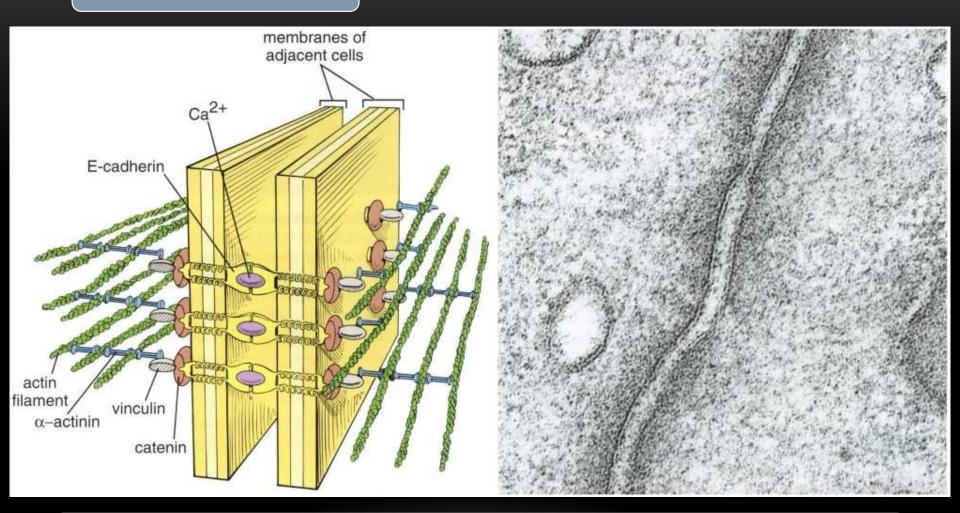
## Anchoring junction (encircles the cell)

AKA adhesion belt, belt junction, or belt desmosome

Located "under" tight junction in epithelial cells

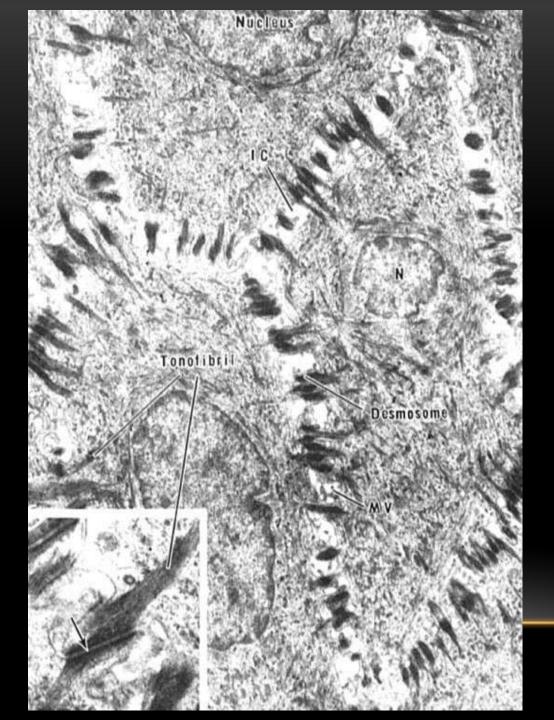
Connected to actin microfilaments that join terminal web

#### **Zonula Adherens**



Cadherin proteins attach to crosslinked actin filaments

Mechanical support - ZA and actin filaments transmit and distribute stress throughout cell and to neighboring cells



#### Desmosomes

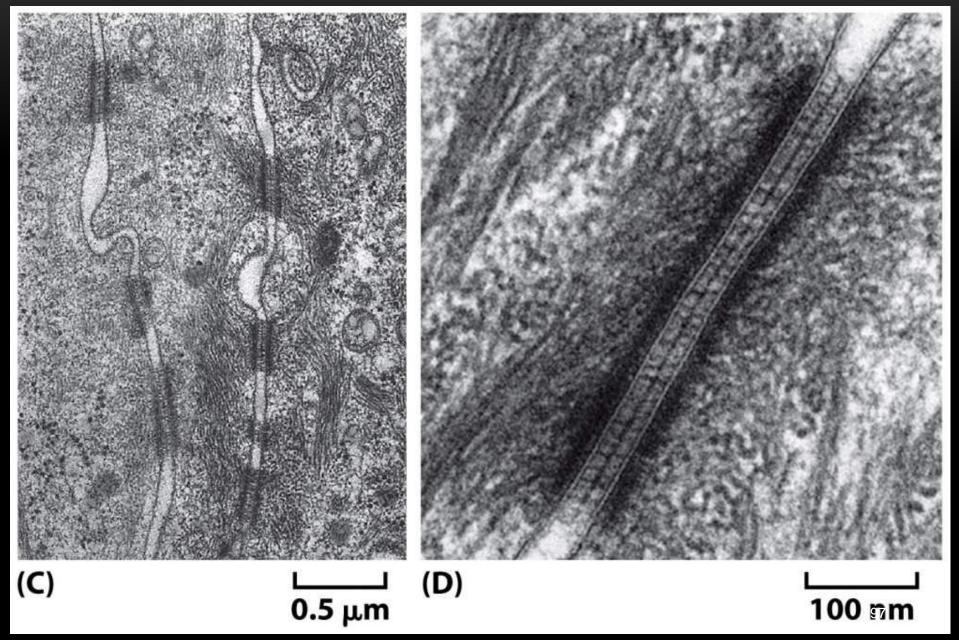
#### Anchoring junctions

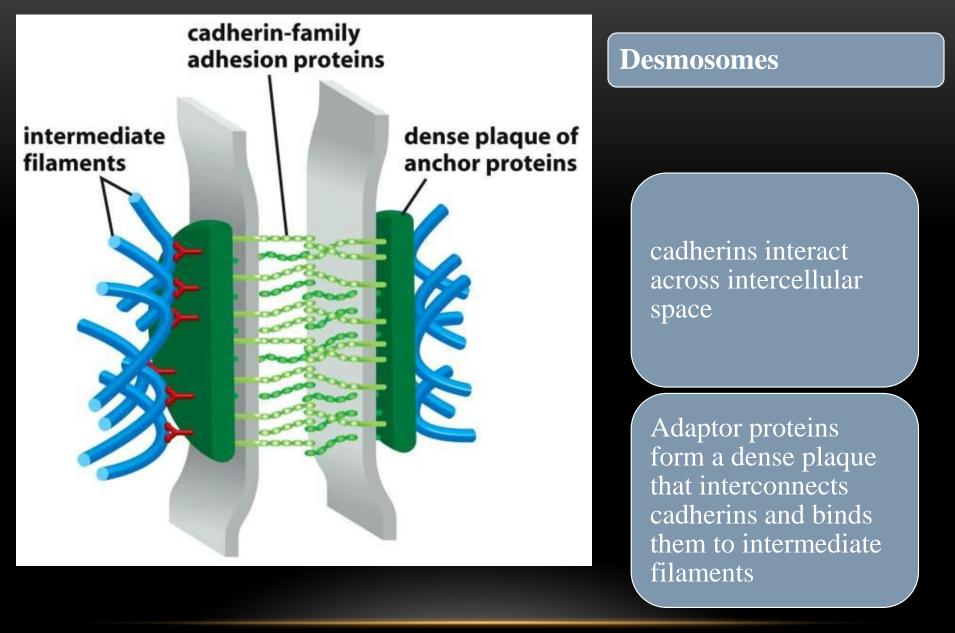
#### Function as "spot welds" to join cells

Located along lateral plasma membranes of columnar epithelial cells or on processes of squamous cells

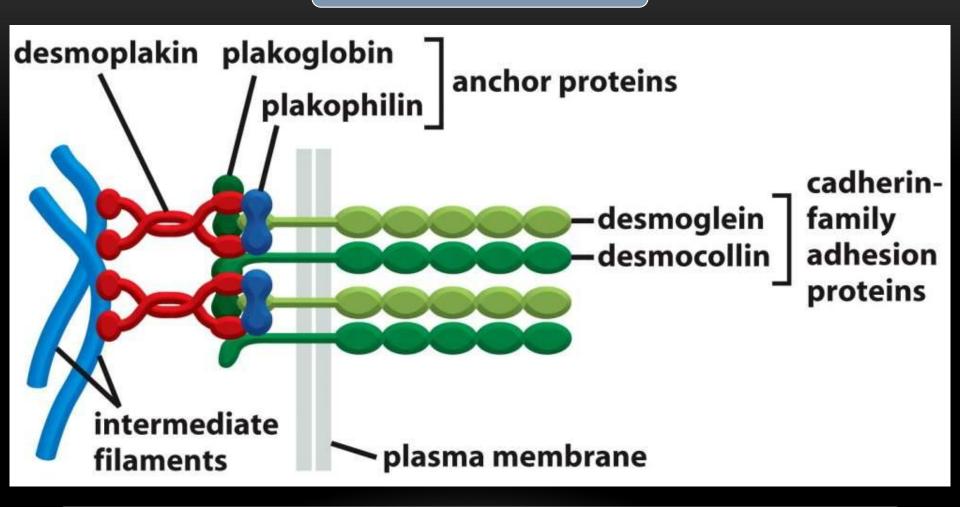
Intermediate filaments associate with plaque proteins in cytoplasm







Desmosomes



Desmoglein and desmocollin are non-classical cadherins

Adaptor proteins such as  $\gamma$ -catenin (plakoglobin) and desmoplakin link cadherins to intermediate filaments

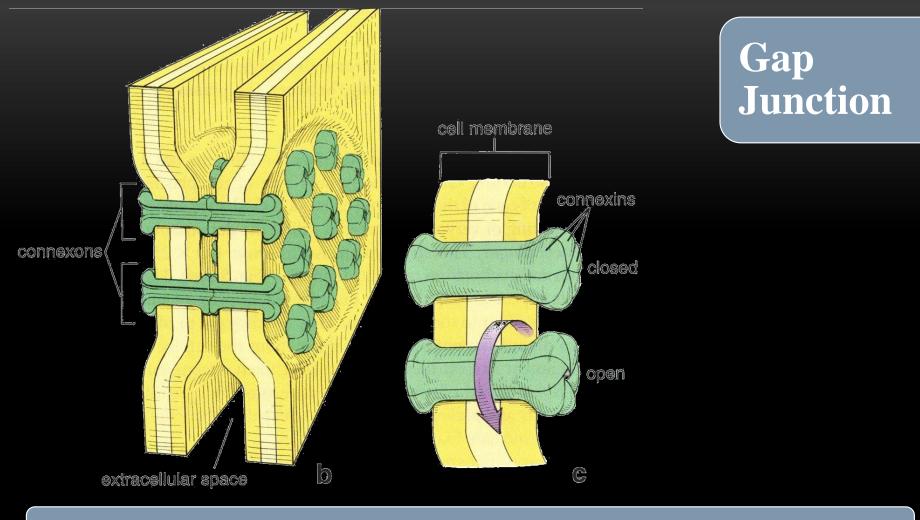


## Gap Junction

#### Channel-forming junction

Named for gap of regular width between cells visualized by TEM

Water-filled junctions transport molecules <1 kDal such as ions, nucleotides (including cAMP), and metabolites



Connexin - protein subunit, six form a hexameric connexon

Connexons - two align to form the gap junction channel

Regulation - elevated calcium concentrations close channel

#### Hemidesmosomes

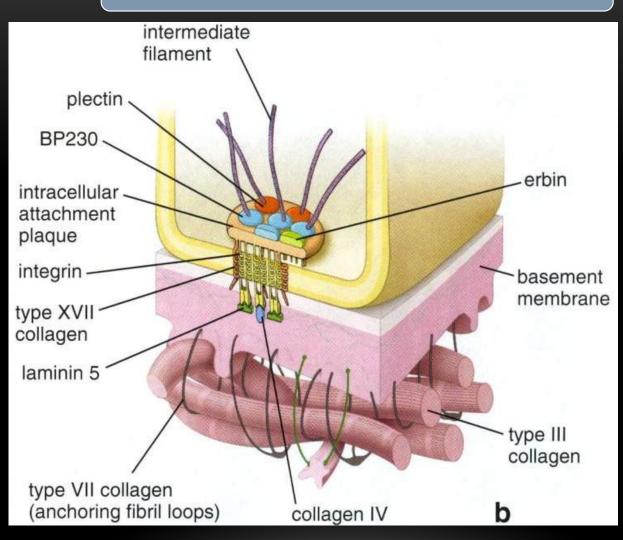


Hemidesmosome - "half-desmosome" in appearance only

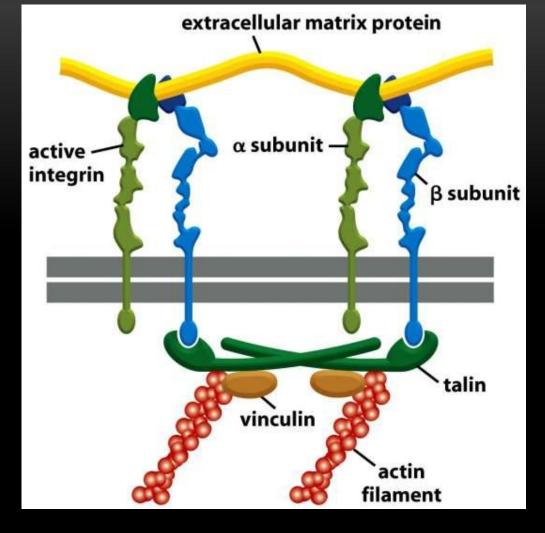
Mediates attachment to basal lamina (extracellular matrix)

Cytoplasmic plaque is attached to cytoskeletal elements

#### Hemidesmosomes



Integrins - membrane protein that "integrates" cell into matrix



#### Integrins

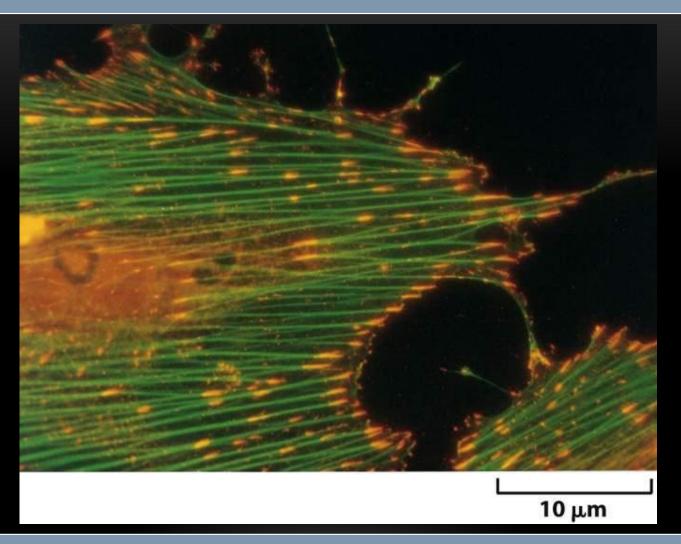
Mediate calcium-independent cell-matrix adhesion

Function as dimers of two membrane proteins ( $\alpha$  and  $\beta$ )

Adaptor proteins link integrins to intermediate filaments in hemidesmosomes or actin filaments in focal adhesions

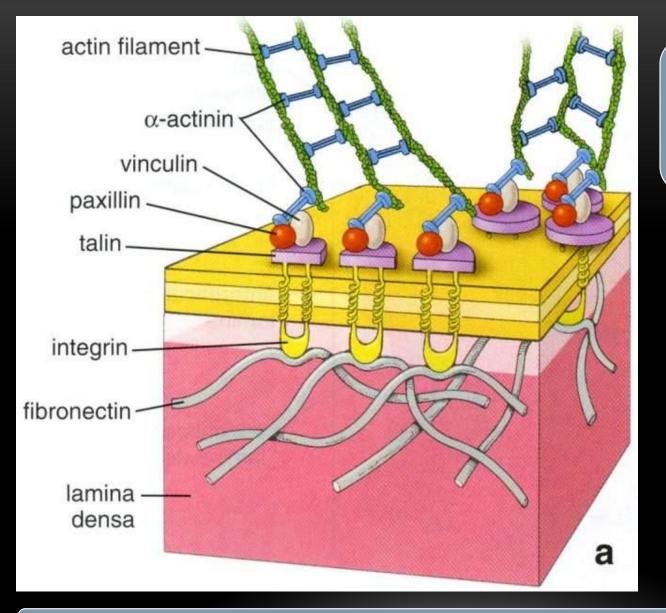
Integrins bind matrix proteins such as laminin or fibronectin

#### **Focal Adhesions**



Anchoring junction (AKA actin-linked cell-matrix adhesion)

Growing fibroblasts form many focal adhesions (orange) that serve as anchoring points for actin filaments (green)



## Focal Adhesions

Fibroblasts attach to extracellular matrix via focal adhesions

Integrins - membrane proteins link actin filaments and matrix

#### **Blistering Disease**



Many mechanisms underlie blistering disorders of the skin

Pemphigus group - autoimmune disease in which autoantibodies target desmogleins present in desmosomes

#### **Pemphigus Histology**



Acantholysis - separation of epidermal keratinocytes (H&E)

# Glandular Epithelium

# **Glands are divided into:**

# Endocrine:

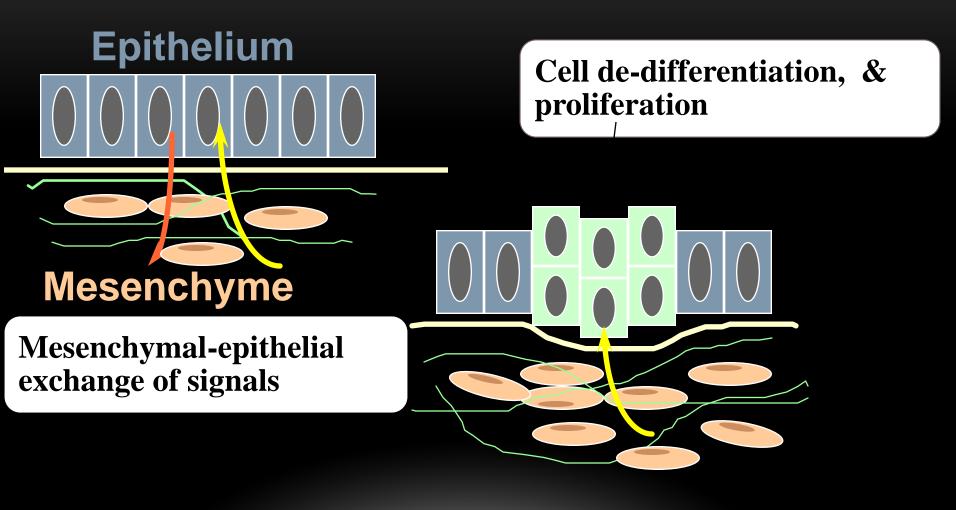
- Unicellular: DNES
- Multicellular: Thyroid, Adrenal

## Exocrine:

- **Unicellular**: Goblet cell
- Multicellular: Parotid, Submandibular, Sublingual

Mixed: Liver, Pancreas, Ovary, Testis.

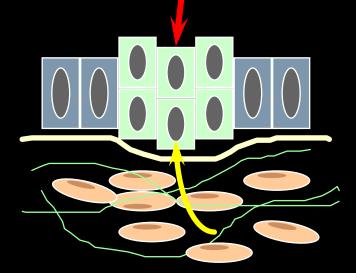
# **Gland Development 1**

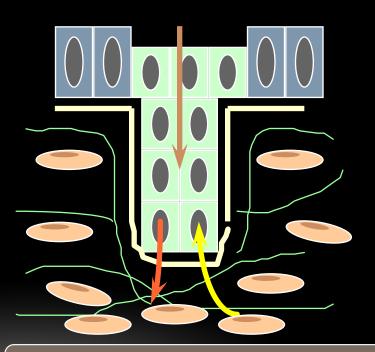


**Gland Development 2** 

#### Cell de-differentiation, & proliferation

Epithelial downgrowth into modified mesenchyme





**Mesenchymal-epithelial interaction** 

### **Gland Development 3**

 $\Box$ 

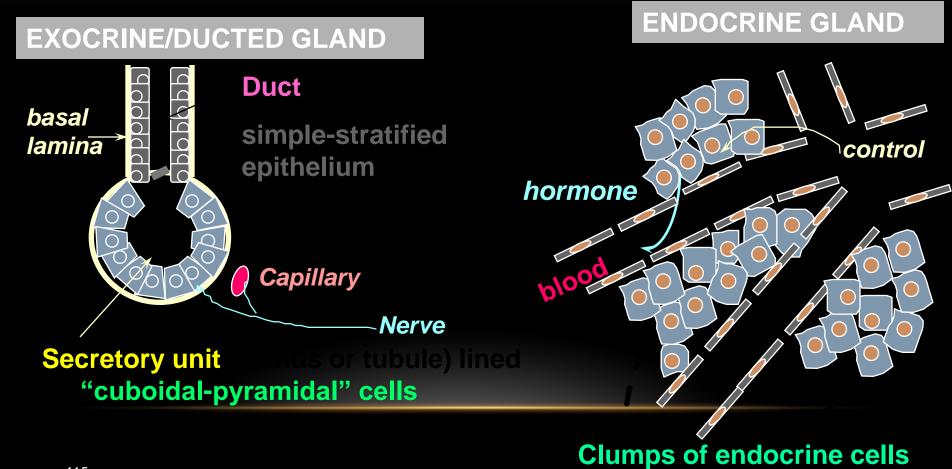
 $\square$ 

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Differentiation into duct & secretory cells

#### **Gland Development 4** qqq -C $\overline{O}$ $\bigcirc$ $\overline{\bigcirc}$ **Construction of lumens** $\cap$ Stroma **Differentiation into** duct & secretory cells Simple alveolar gland

Although cells i GLANDULAR EPITHELIA crete, they are limited in number. To get more secreting power, and sometimes to focus it differently, e.g. to interact with blood, rather than dump into a principal tube, epithelial cells can build *glands* 



## EXOCRINE GLANDS.. 1:

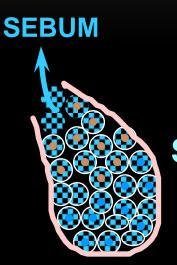
# <u>Classified according to the mode of secretion:</u>

# Merocrine (eccrine): Apocrine: Holocrine:

# **MODES OF SECRETORY RELEASE**

## MEROCRINE / ECCRINE

hormone



secretion released by exocytosis, with *no loss* of cytoplasm

Sebaceous gland

e.g., by endocrine cells

secretion released, filling a *dead* cell

<u>HOLOCRINE</u>

released by exocytosis, with *a little loss of cytoplasm* 

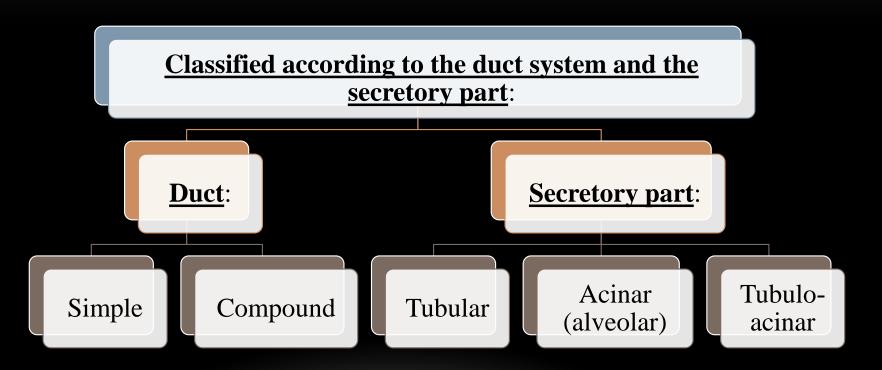


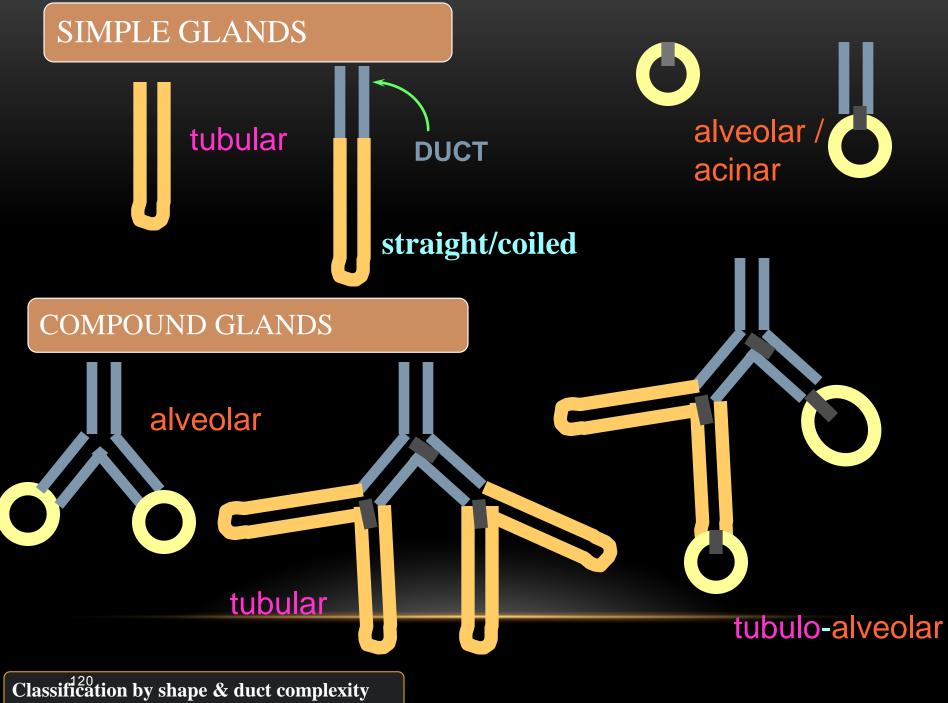
# **Exocrine Glands.. 2:**

# <u>Classified according to</u> <u>nature of secretion</u>:

- <u>Serous</u>:
- <u>Mucous</u>:
- <u>Mixed</u>:

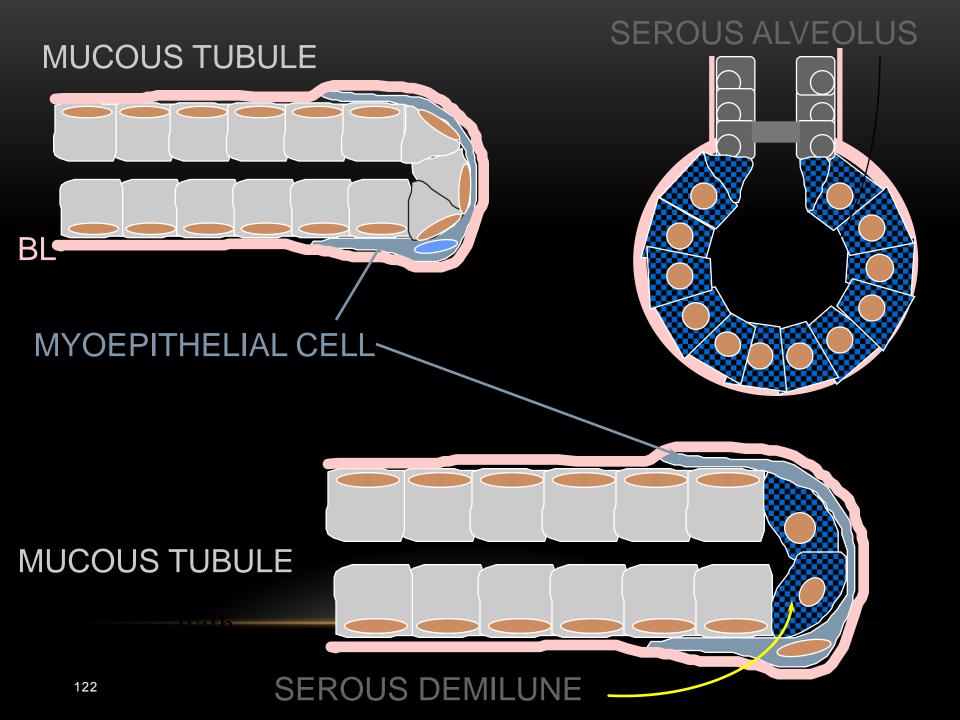
# **Exocrine Glands.. 3:**



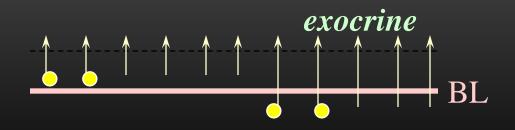


# **Glandular Epithelia : Products & Roles**

Extra mucus	Airway glands, Duodenal & Salivary glands
Extra defense	Airway glands
Digestion	Gastric glands, Pancreas
Blood processing	Liver
Hormones	Endocrine glands
Milk	Mammary glands
Sweat	Sweat glands
Grease	Sebaceous glands
Genital glands	Special genito-urinary functions 121



Deliver all along a surface Single cells



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MUCUS-SECRETING GOBLET CELL

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0

*muco-ciliary escalator* rids airway of particles

#### Deliver all along a surface

#### Surface goblet cells

**Colon** cells

Simple straight tubules, with - SIMPLE TUBJLAR

# **GUT PARTS**

**MUSCULARIS** smooth muscle

STREET, STREET

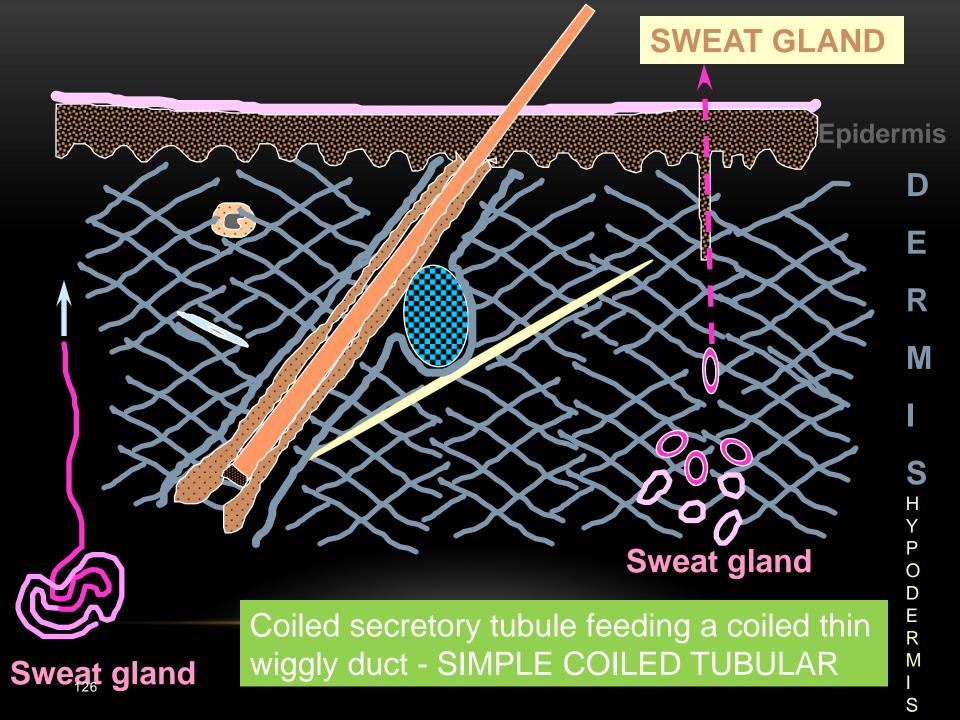
# suspensory **MESENTERY** with blood vessels

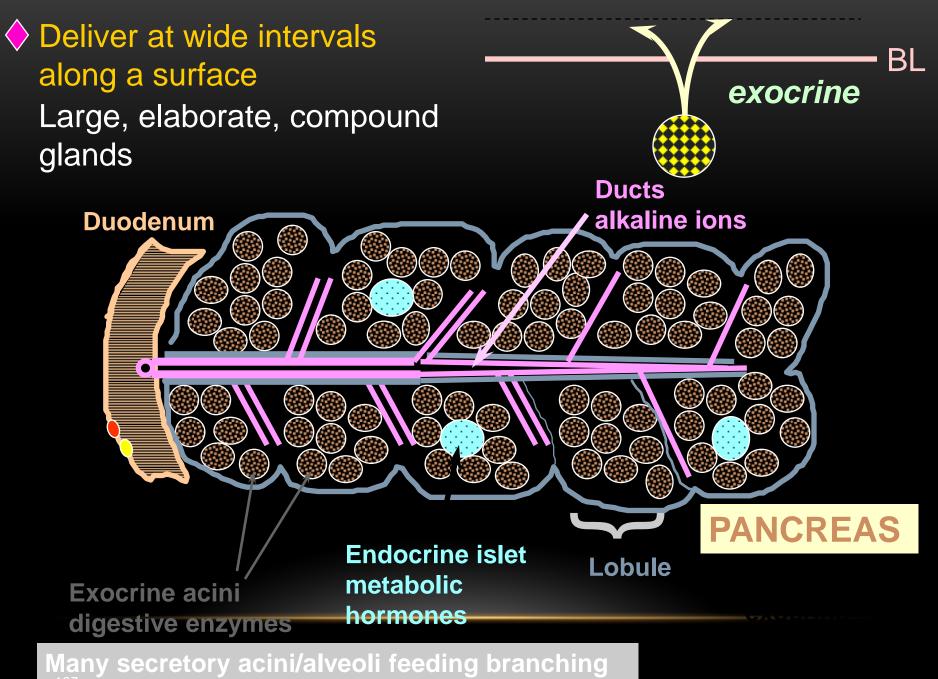
confrective tissue

simple columnar epithelium GLANDS

covering **SEROSA** with simple squamous epithelium







duct system - COMPOUND ALVEOLAR

# Duodenal **EXOCRINE PANCREAS** Ducts 1

#### papilla

## Principal duct

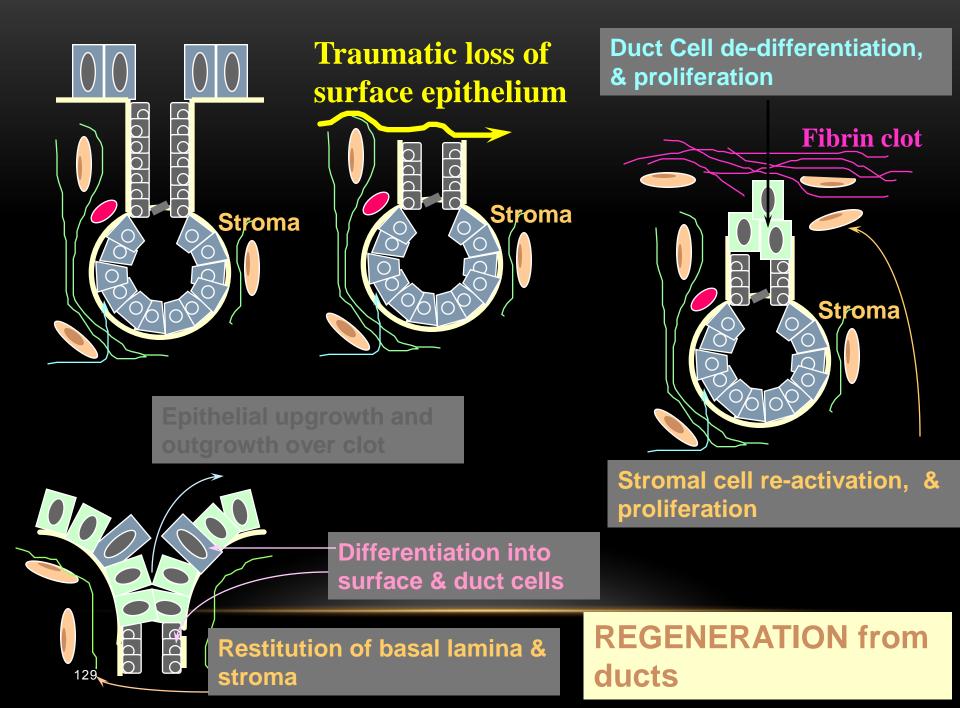
## Interlobular duct

#### Intralobular ducts

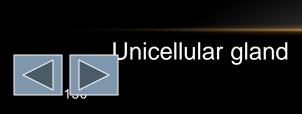
#### Intercalated ducts

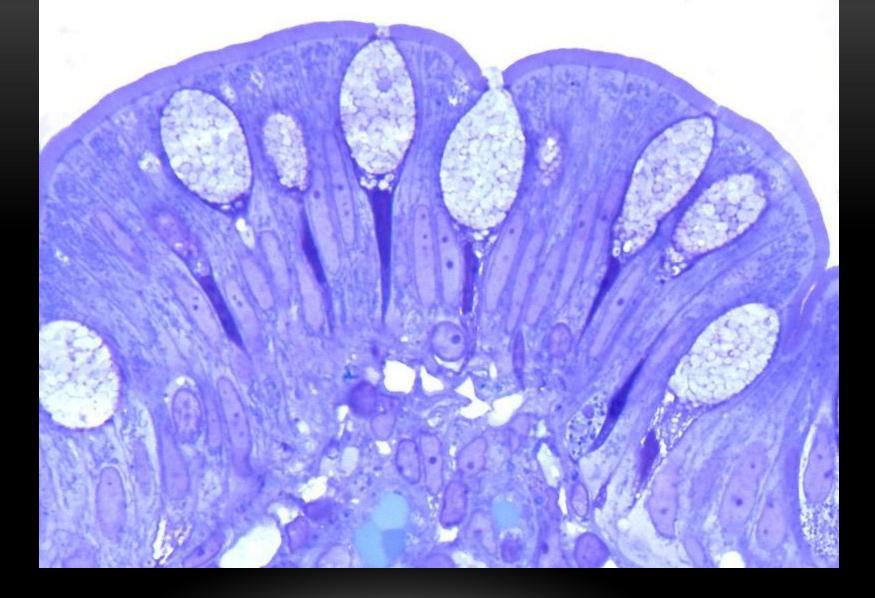
**Exocrine** acini

Lobule



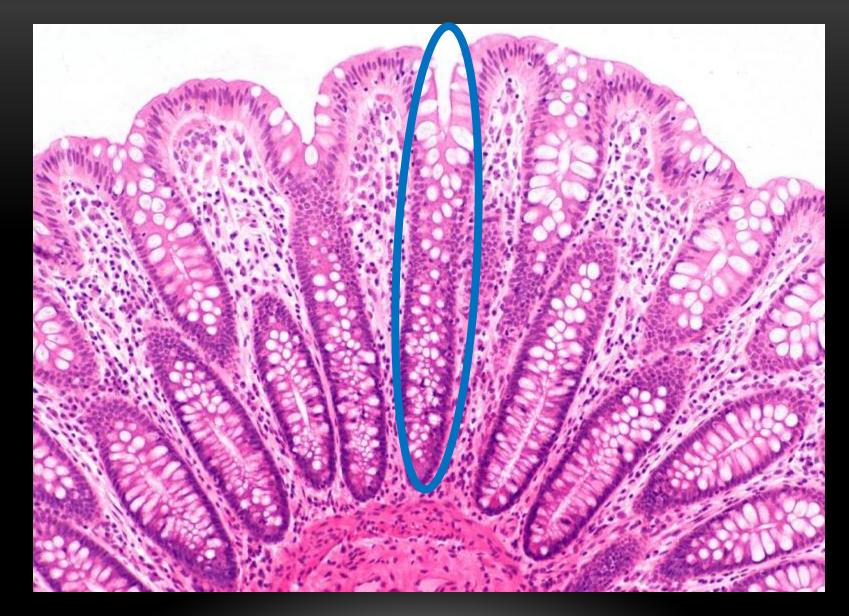






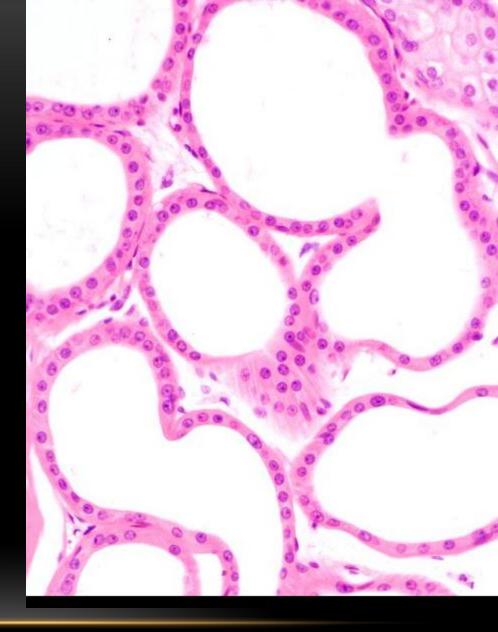
Unicellular gland



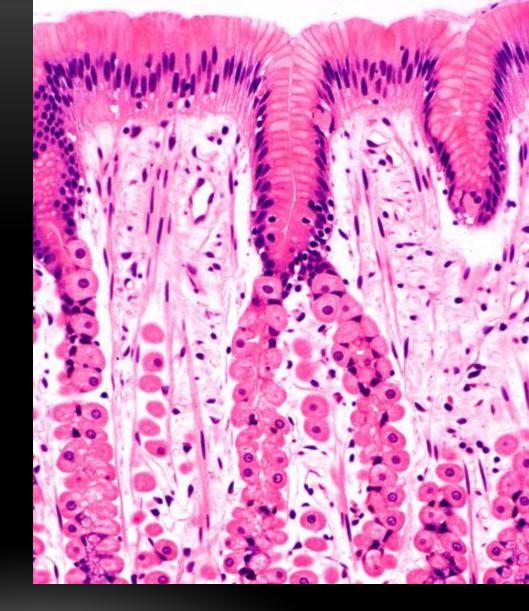


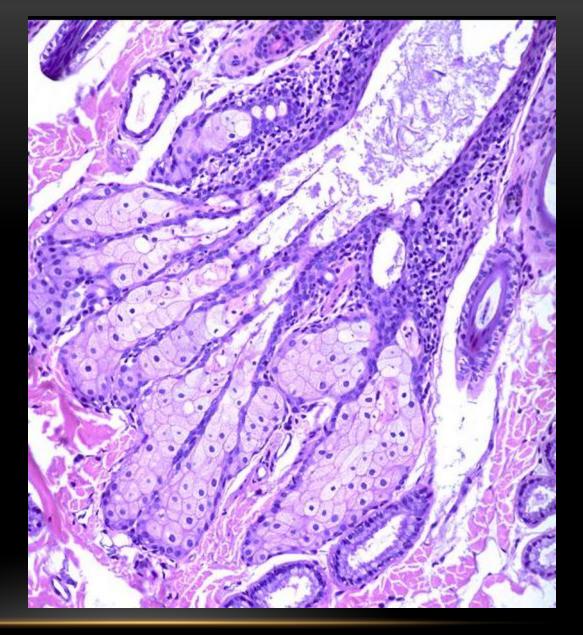
Simple tubular glands

#### Simple coiled tubular gland

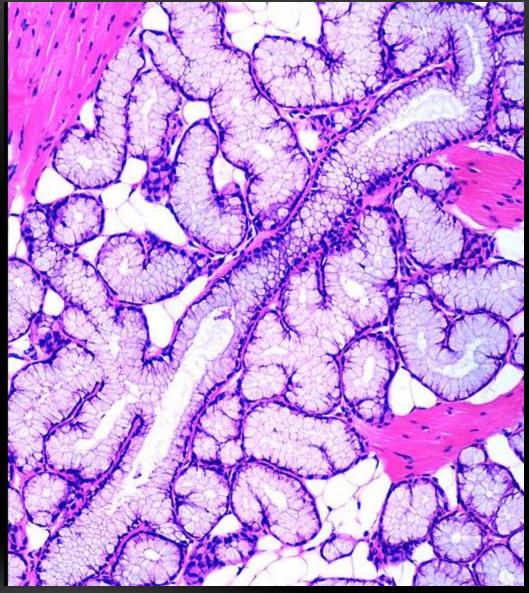


#### Simple branched tubular

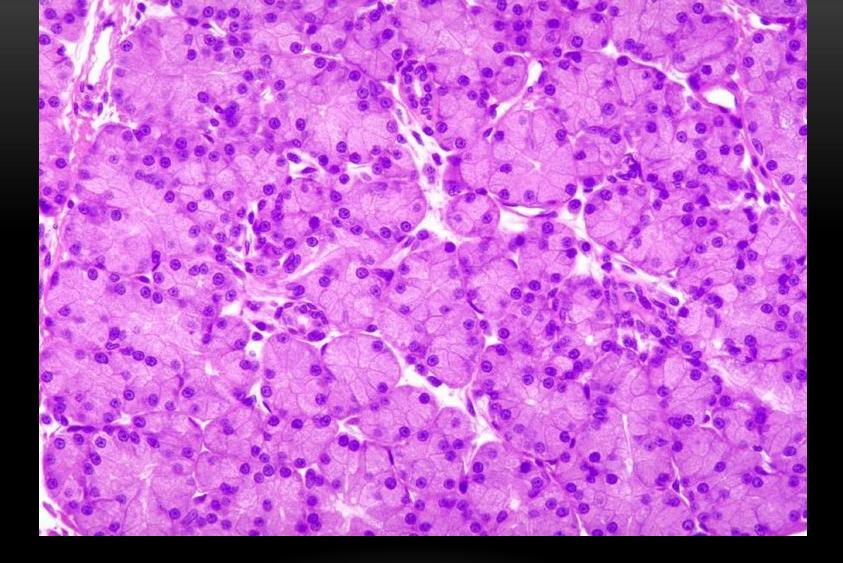




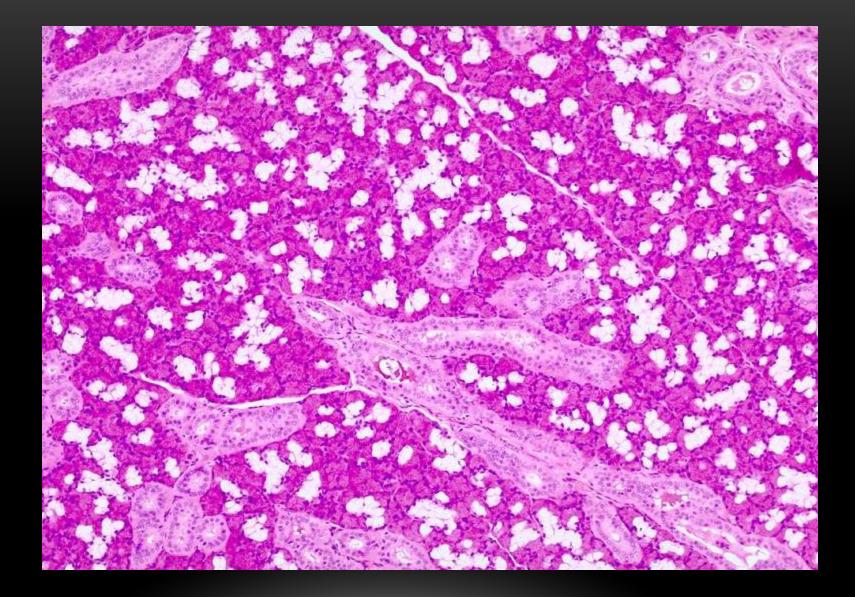
Simple alveolar glands



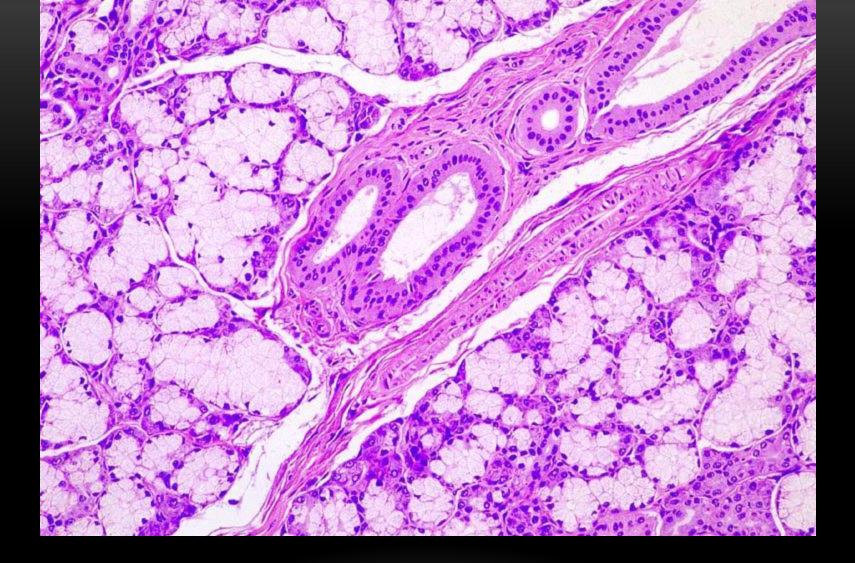
#### Compound tubular glands



Serous gland- parotid



Seromucous gland



Mucus gland



Apocrine gland

