Epihelial tissue

CHARACTERISTIC FEATURES OF EPITHELIAL CELLS Basement Membranes Intercellular Adhesion & Other Junctions SPECIALIZATIONS OF THE APICAL CELL SURFACE Microvilli Stereocilia Cilia	73 74 75 78 79 81 81	TYPES OF EPITHELIA Covering or Lining Epithelia Secretory Epithelia & Glands TRANSPORT ACROSS EPITHELIA RENEWAL OF EPITHELIAL CELLS SUMMARY OF KEY POINTS	81 81 86 90 91
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Human body:

- 1. Epithelial tissue
- 2. Connective tissue
- 3. Mascular tissue
- 4. Nervous tissue

TABLE 4-1	Main characteristics of the four basic types of tissues.				
Tissue	Cells	Extracellular Matrix	Main Functions		
Nervous	Elongated cells with extremely fine processes	Very small amount	Transmission of nerve impulses		
Epithelial	Aggregated polyhedral cells	Small amount	Lining of surface or body cavities; glandular secretion		
Muscle	Elongated contractile cells	Moderate amount	Strong contraction; body movements		
Connective	Several types of fixed and wandering cells	Abundant amount	Support and protection of tissues/ organs		

Organs:

- Parenchyma
- Stroma

Epihelial tissue

3

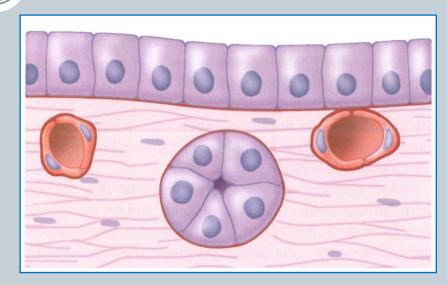
- Covering, lining & protecting surface (epidermis)
- Absorbtion (the intestinal lining)
- Secretion (parenchymal cell of glands)

Specific cells:

- Myoepithelial cells (contractile)
- Taste bubs or olfactory epithelium (specialized sensory cells)

Characteristic features of epithelial cells

- 4
- Columnar cell (elliptic or oval nucleus)
- Cuboidal cells (spherical nucleus)
- Squamous cell (flattened nucleus)



- No blood vessle
- Lamina propria in digestive, respiratory & urinary systems
- Papillae (in skin & tongue)
- PolarityBasal poleApical poleLateral surface

Basement membrane

- Extra cellular materials
- o Glycoprotein & ...

By TEM:

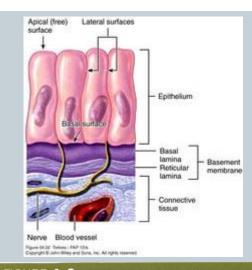
- 1. Basal lamina (electron dens 20-100 nm)
- 2. Reticular lamina (diffuse & fibrous)

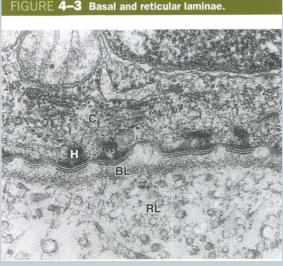
Basal lamina macromolecules

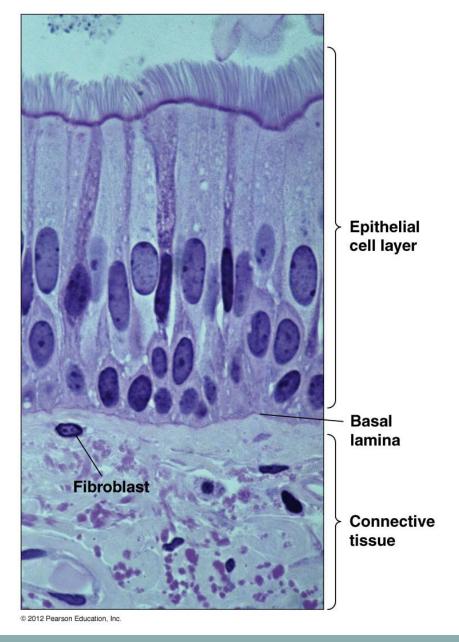
- 1. Laminin (glycoprotein)
- 2. type IV Collagen (3 chain poly peptide)
- 3. Entactin/nidogen(adhesive glycoprotein)
- 4. Perlecan (proteoglycan that bind collagen IV & laminin)

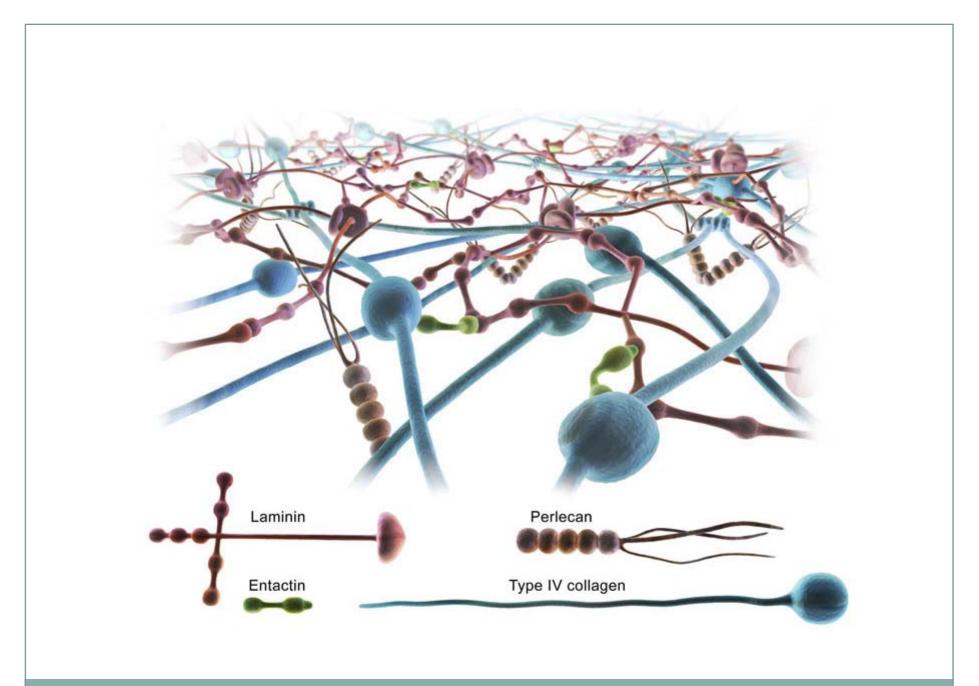
Reticular lamina macromolecules

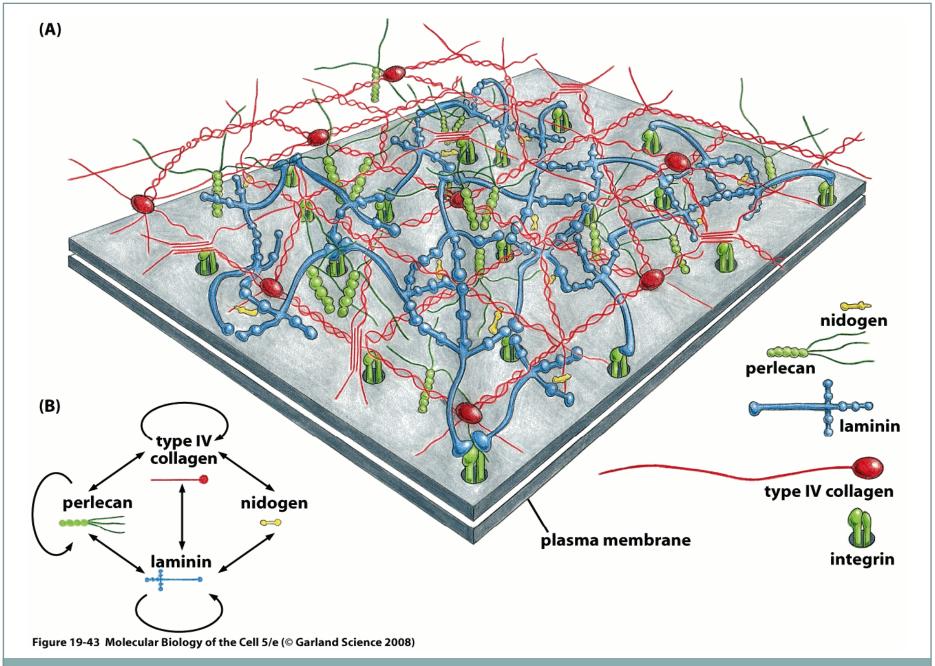
- 1. Type III collagen
- 2. Type VII collagen (bind type III Col. To basal lamina)

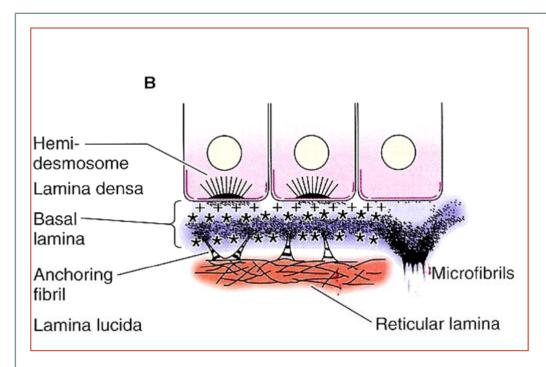


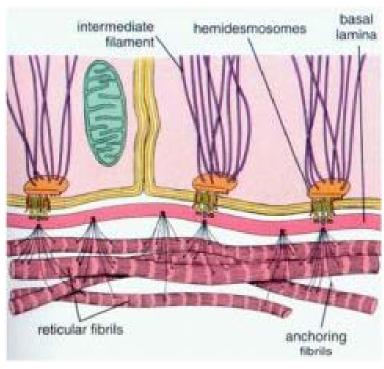










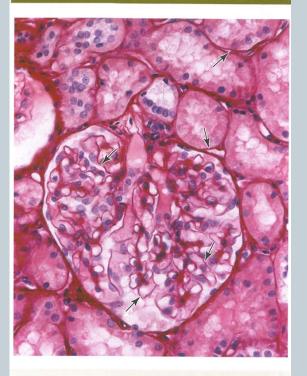


Basement membrane function

10

- Structural support
- Polarity
- Attachment
- Filtration
- Mitogenic factor concentration
- Scaffold for repair & regeneration
- Protein organization in plasma membrane
- Cell-cell interaction (migration)

FIGURE 4-2 Basement membranes.



This section of kidney shows the well-stained basement membranes (arrows) of epithelia forming structures within the large, round renal glomerulus and its surrounding tubules. In kidney glomeruli the basement membrane, besides having a supporting function, has a highly developed role as a filter that is key to renal function. X100. Picrosiriushematoxylin (PSH).

Intercellular adhesion & other junction

- 1. Tight junction(occluding j.)
- 2. Adherent junction (anchoring j.)
- 3. Gap junction (communicating j.)
- Zonula occludense (cloudin & occludin pr.)
- Zonula adherens (cadherin)
 Catenin & actin (terminal web)
- Desmosome or macula adherens (anchoring j.)
 Desmoglein & desmocollin (cadherin)
 Desmoplakin & plakoglobin
 Cytokeratin (desmin & vimentin)
- Gap junction (connexin make connexon)
- Hemidesmosome integrin bind to laminin & col. IV

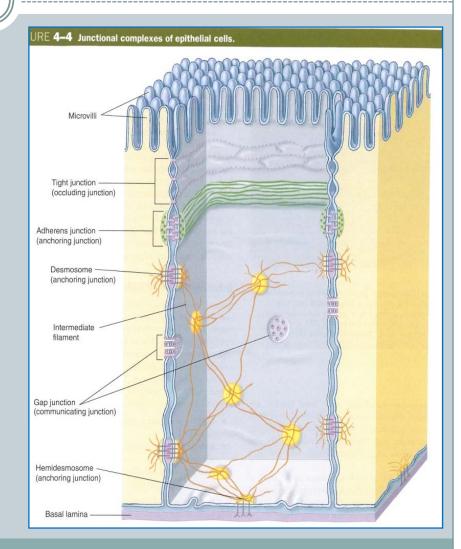
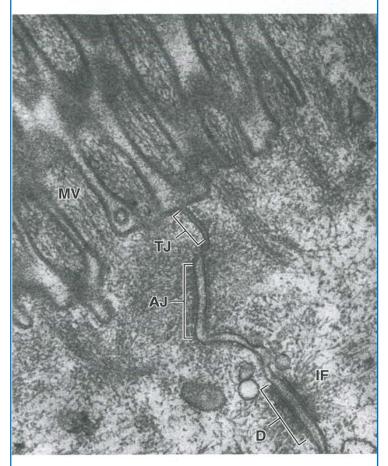


FIGURE 4-5 Epithelial cell junctional complex.



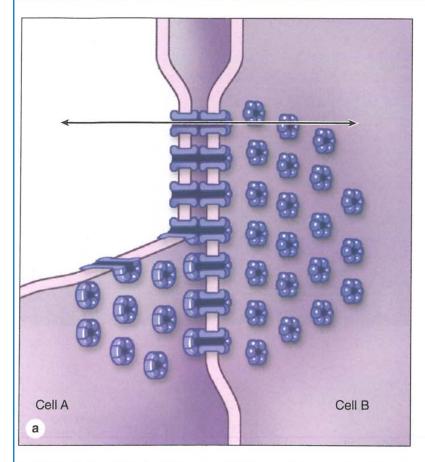
Ultrastructural view of the apical region near microvilli (MV) of two epithelial cells, revealing a junctional complex with a tight junction (TJ) or zonula occludens, an adherent junction (AJ) or zonula adherens, and a desmosome (D) associated with intermediate filaments (IF). The functions and major protein components of these junction types are summarized in Table 4–2. \times 195,000.

FIGURE **4–6** View of a tight junction after cryofracture.



Just below the apical microvilli (MV) of this epithelial cell, a cryofracture plane splitting fused cell membranes reveals the fused strands of transmembrane proteins forming the tight junction (zonula occludens). X100,000.

FIGURE 4-7 Gap junctions.





(a) A diagram of a gap junction shows the structural elements that allow the exchange of nutrients and signal molecules between cells without loss of material into the intercellular space. The communicating channels are formed by pairs of abutting particles (connexons), which are in turn each composed of six protein subunits (connexins) that span the lipid

bilayer of each cell membrane. The channel formed by paired connexons (arrow) is about 1.5 nm in diameter, limiting the size of transmitted molecules. (b) A cryofracture preparation of a gap junction, showing the patch of aggregated transmembrane protein complexes, the connexons. X150,000.

TABLE **4–2**

Epithelial cell junctions, their major structural features and functions, and medical significance.

Junction	Tight Junction (Zonula Occludens)	Adherent Junction (Zonula Adherens)	Desmosome (Macula Adherens)	Hemidesmosome	Gap Junction (Nexus)
Major transmembrane link proteins	Occludins, claudins, ZO proteins	E-cadherin, catenin complexes	Cadherin family proteins (desmogleins, desmocollin)	Integrins	Connexin
Cytoskeletal components	Actin filaments	Actin filaments	Intermediate filaments (keratins)	Intermediate filaments	None
Major functions	Seals adjacent cells to one another, controlling passage of molecules between them; separates apical and basolateral membrane domains	Provides points linking the cytoskeletons of adjacent cells; strengthens and stabilizes nearby tight junctions	Provides points of strong intermediate filament coupling between adjacent cells, strengthening the tissue	Anchors cytoskeleton to the basal lamina	Allows direct transfer of small molecules and ions from one cell to another
Medical significance	Defects in occludins may compromise the fetal blood-brain barrier, leading to severe neurologic disorders	Loss of E-cadherin in epithelial cell tumors (carcinomas) promotes tumor invasion and the shift to malignancy	Autoimmunity against desmoglein I leads to dyshesive skin disorders characterized by reduced cohesion of epidermal cells	Mutations in the integrin-β4 gene are linked to some types of epidermolysis bullosa, a skin blistering disorder	Mutations in various connexin genes have been linked to certain types of deafness and peripheral neuropathy

>> MEDICAL APPLICATION

Various **blistering** (**bullous**) **diseases**, such as pemphigus vulgaris, involving the epidermis or stratified squamous epithelia of the oral mucosa, are due to abnormal desmosome function caused by autoimmune reactions against specific desmogleins that reduce cell-to-cell adhesion. Similar disorders arise with genetic mutations for various junctional proteins.

>> MEDICAL APPLICATION

Proteins of the **zonula occludens** provide the targets for certain common bacteria of medical importance. The enterotoxin secreted by *Clostridium perfringens*, which causes "food poisoning," binds claudin molecules of intestinal cells, prevents insertion of these proteins during maintenance of tight junctions, and causes loss of tissue fluid into the intestinal lumen via the paracellular pathway.

Similarly, *Helicobacter pylori*, which is important in the etiology of **gastric ulcers**, binds the extracellular domains of tight-junction proteins in cells of the stomach and inserts a protein into these cells, which targets ZO-1 and disrupts signaling from the junction.

Specializations of the apical cell surface

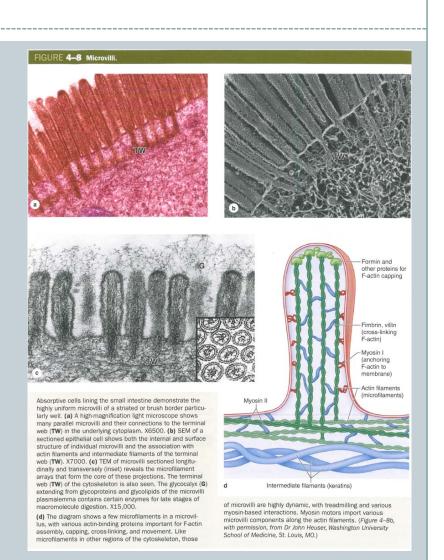
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Microvilli

- Brush or striated border
- 1 μ length & 0.1 μ
- Enhance surface (20-30)
- Thick glycolalix (enzymes)

>> MEDICAL APPLICATION

Celiac disease, also called gluten-sensitive enteropathy or sprue, is a disorder of the small intestine in which one of the first pathologic changes is loss of the microvilli brush border of the absorptive cells. This is caused by an immune reaction against the wheat protein gluten during its digestion, which produces diffuse enteritis (intestinal inflammation), changes to the epithelial cells leading to malabsorption, and eventually to pathologic changes in the intestinal wall. The malabsorption problems and structural changes are reversible when gluten is removed from the diet.



Stereocilia



- Epididymis
- Proximal ductus deferens
- Increase cell surface area
- Increse absorption
- Inner ear sensory cells
- motion detecting function
- Longer than microvilli & less motile

In terminal web

FIGURE 4-9 Stereocilia.



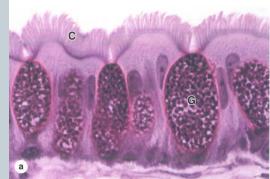
At the apical ends of the tall epithelial cells lining organs such as the epididymis (shown here) are numerous very long stereocilia, which increase the surface area available for absorption. Stereocilia are much longer than microvilli and often have distal branching. X400. H&E.

Cilia

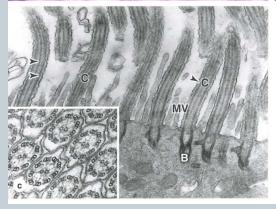


- Longer than microvilli
- Microtubule
- Primary cilia
- Motile cilia (epithelial cells)
- 9+2 doublet structure
- 9 triplets in basal body
- Dynein & kinesin
- ATP
- Flagellum



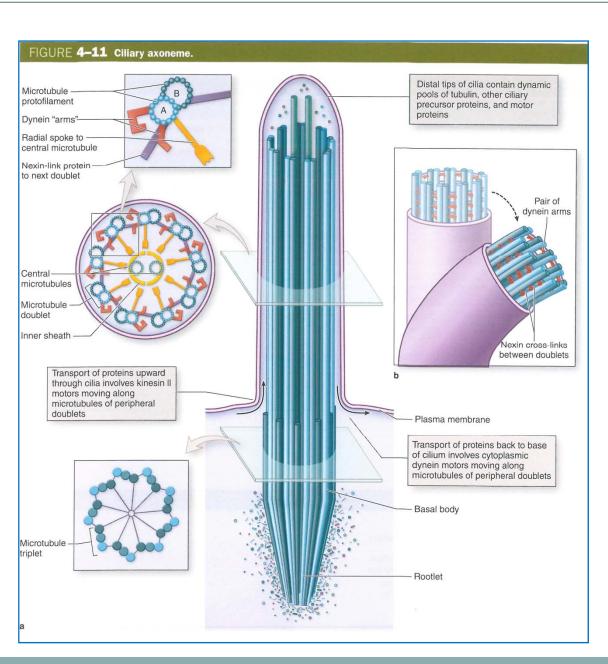






Epithelial cells lining the respiratory tract have many very well-developed cilia. (a) By light microscopy cilia (C) on the columnar cells appear as a wave of long projections, interrupted by nonciliated, mucus-secreting goblet cells (G). X400. Toluidine blue. (b) SEM of the apical surfaces of this epithelium shows the density of the cilia (C) and the scattered goblet cells (G). X300.

(c) TEM of cilia (C) sectioned longitudinally reveals the central and peripheral microtubules of the axonemes, with cross sections (inset) clearly showing the 9+2 array of the microtubule doublets. At the base of each cilium is a basal body (B) anchoring the axoneme to the apical cytoplasm. Much shorter microvilli (MV) can be seen between the cilia. X59,000. Inset: X80,000.



Epithelia types

20)

- Covering or lining epithelium
- Secretory or glandular epithelium

Covering or lining epithelium

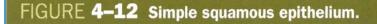
- ☐ Simple epithelia
- Stratified epithelia
- Pseudostratified columnar epithelium

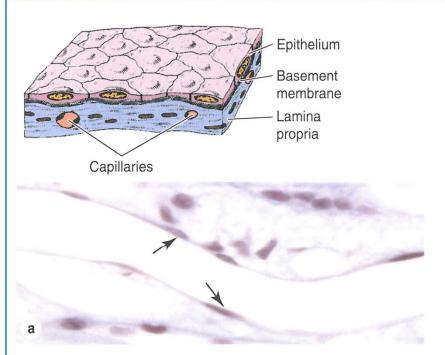
☐ Simple epithelia

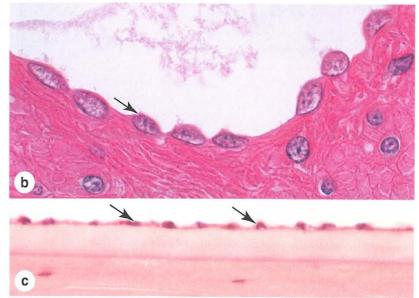
Squamous

Cuboidal

Columnar







This is a single layer of thin cells, in which the **cell nuclei** (arrows) are the thickest and most visible structures. Simple epithelia are typically specialized as lining of vessels and cavities, where they regulate passage of substances into

the underlying tissue. The thin cells often exhibit transcytosis. Examples shown here are those lining the thin renal loops of Henle (a), covering the outer wall of the intestine (b), and lining the inner surface of the cornea (c). a, c X400; b X600. H&E.

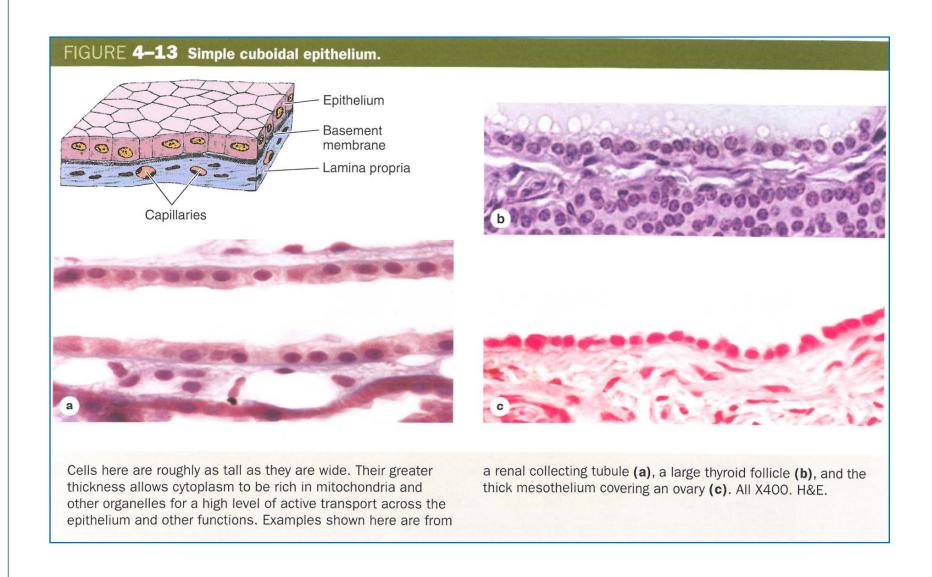
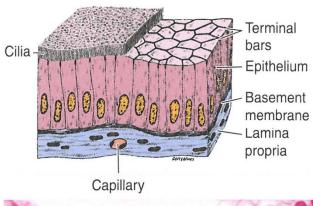
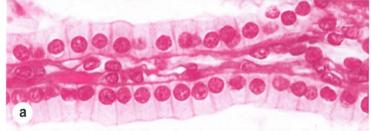
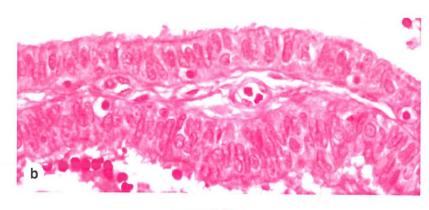
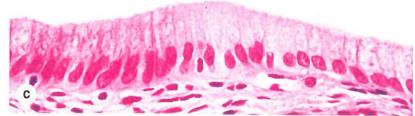


FIGURE 4-14 Simple columnar epithelium.









Cells here are always taller than they are wide, with apical cilia or microvilli, and are often specialized for absorption. Complexes of tight and adherent junctions, sometimes called "terminal bars" in light microscopic images, are present at the

apical ends of cells. The examples shown here are from a renal collecting duct (a), the oviduct lining, with both secretory and ciliated cells (b), and the lining of the gall bladder (c). All X400. H&E.

Stratified epithelia

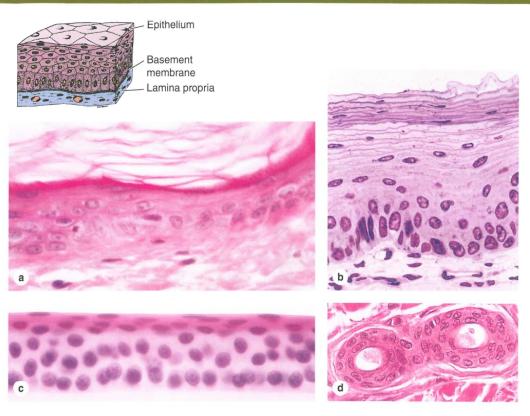
Squamous

- 1. Keratinized (skin)
- 2. Nonkeratinized (mouth, esophagus, vagina)

Cuboidal (exceretory ducts of salivary & sweet glands)
Columnar (conjunctiva lining eyelids, protective & mucos secretion)
Transitional or urothelium (umbrella cells)

☐ Pseudostratified columnar epithelium

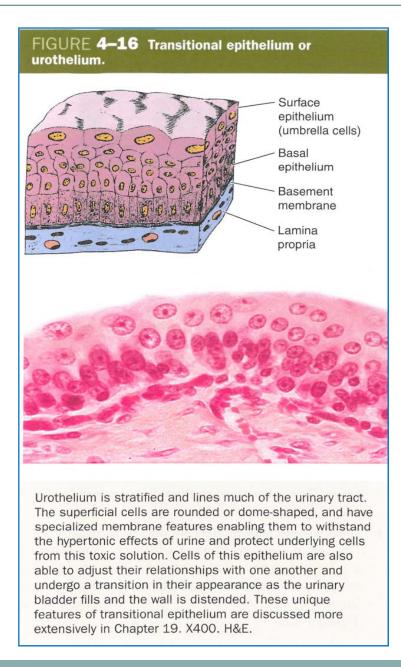
FIGURE 4-15 Stratified epithelium.



Stratified squamous epithelia usually have protective functions: protection against easy invasion of underlying tissue by microorganisms and protection against water loss. These functions are particularly important in the epidermis (a) in which differentiating cells become **keratinized**, ie, filled with keratin and other substances, eventually lose their nuclei and organelles, and form superficial layers flattened squames that impede water loss. Keratinized cells are sloughed off and replaced by new cells from more basal layers, which are discussed fully with the skin in Chapter 18.

Nonkeratinized epithelia occur in many organs, such as the esophageal lining **(b)** or outer covering of the cornea **(c)**. Here cells accumulate much less keratin and retain their nuclei but still provide protection against microorganisms.

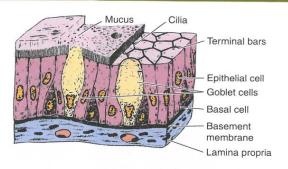
Stratified cuboidal or columnar epithelia are fairly rare but occur in excretory ducts of certain glands, such as sweat glands (d) where the double layer of cells allows additional functions. All X400; (b) PT, (a, c, and d) H&E.



Pseudostratified columnar epithelium



FIGURE 4-17 Pseudostratified epithelium.





Cells of pseudostratified epithelia appear to be in several layers, but their basal ends all rest on the basement membrane. The pseudostratified columnar epithelium of the upper respiratory tract shown here contains many ciliated cells, as well as other cells with their nuclei at different levels. X400. H&E.

>> MEDICAL APPLICATION

In **chronic bronchitis**, common among habitual smokers, the number of goblet cells in the lining of airways in the lungs often increases greatly. This leads to excessive mucus production in areas where there are too few ciliated cells for its rapid removal and contributes to obstruction of the airways. The ciliated pseudostratified epithelium lining the bronchi of smokers can also be transformed into stratified squamous epithelium by metaplasia.

TABLE 4-3

Common types of covering epithelia.

Major Feature	Cell Form	Examples of Distribution	Main Function
Simple (one layer of cells)	Squamous	Lining of vessels (endothelium); Serous lining of cavities: pericardium, pleura, peritoneum (mesothelium)	Facilitates the movement of the viscera (mesothelium), active transport by pinocytosis (mesothelium and endothelium), secretion of biologically active molecules (mesothelium)
	Cuboidal	Covering the ovary, thyroid	Covering, secretion
	Columnar	Lining of intestine, gallbladder	Protection, lubrication, absorption, secretion
Pseudostratified (layers of cells with nuclei at different levels; not all cells reach surface but all adhere to basal lamina)		Lining of trachea, bronchi, nasal cavity	Protection, secretion; cilia-mediated transport of particles trapped in mucus out of the air passages
Stratified (two or more layers of cells)	Squamous keratinized (dry)	Epidermis	Protection; prevents water loss
	Squamous nonkeratinized (moist)	Mouth, esophagus, larynx, vagina, anal canal	Protection, secretion; prevents water loss
	Cuboidal	Sweat glands, developing ovarian follicles	Protection, secretion
	Transitional	Bladder, ureters, renal calyces	Protection, distensibility
	Columnar	Conjunctiva	Protection

Secretory epithelia & glands

29)

Secretory vesicle

- Protein (pancreas)
- Lipid (adrenal & sebaceous glands)
- Carbohydrate and pr. Complex (salivary glands)
- All of mentioned (mammary glands)
- Water & electrolyte (sweet glands)

• Unicellular glands Goblet cell



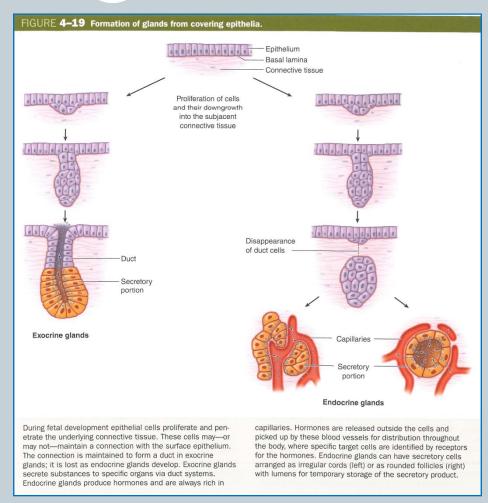
The simple columnar epithelium lining the large intestine shows many isolated goblet cells secreting mucus into the lumen. (a) With a stain for the oligosaccharide components of mucin glycoproteins, the cytoplasmic secretory granules of two goblet cells and secreted mucus are stained purple. X600. PAS-PT. (b) As shown ultrastructurally, goblet cells always have

basal nuclei surrounded by RER (**R**), a large Golgi complex (**G**), and abundant apical cytoplasm filled with large secretory granules (**SG**). After exocytosis mucin components are hydrated and become mucus. A brush border of microvilli (**M**) is seen on neighboring columnar cells. X17,000.

Gland formation

(30)

- Exocrine glands
- Endocrine glandsSecretory portions & ducts



Exocrine Glands

(31)

• Glands & ducts:

Ducts

- 1. Simple
- 2. Compound

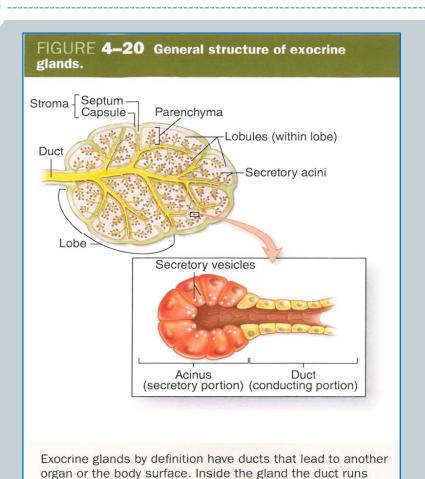
Tubular

Acinar

Tubuloacinar

Secretory portion

- Tubular (short or long & coiled)
- coiled
- Acinar



through the connective tissue of septa and branches repeatedly, until its smallest branches end in the secretory portions

of the gland.

TABLE 4-4

Structural classes of exocrine glands, features of each class, and examples.

SIMPLE Glands (Ducts Do Not Branch)

Class	Simple Tubular	Branched Tubular	Coiled Tubular	Acinar (or Alveolar)	Branched Acinar
	Duct-Secretory portion				THE PARTY OF THE P
Features	Elongated secretory portion; duct usually short or absent	Several long secretory parts joining to drain into 1 duct	Secretory portion is very long and coiled	Rounded, saclike secretory portion	Multiple saclike secretory parts entering the same duct
Example	Mucous glands of colon; intestinal glands or crypts (of Lieberkühn)	Glands in the uterus and stomach	Sweat glands	Small mucous glands along the urethra	Sebaceous glands of the skin

COMPOUND Glands (Ducts from Several Secretory Units Converge into Larger Ducts) Tubular Acinar (Alveolar) Tubuloacinar Class Secretor portions Several elongated, coiled secretory units Several saclike secretory units with Features Ducts of both tubular and and their ducts converge to form larger small ducts converge at a larger acinar secretory units ducts duct converge at larger ducts Submucosal mucous glands (of Brunner) Exocrine pancreas Examples Salivary glands in the duodenum

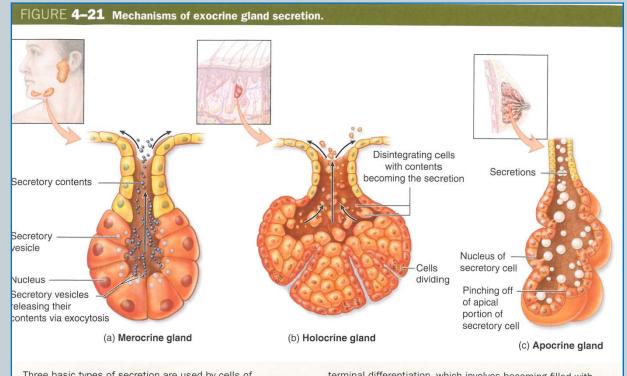
Secretion mechanism

34

- Merocrine secretion
- Apocrine secretion
- Holocrine secretion

>> MEDICAL APPLICATION

The holocrine sebaceous glands are the primary structure involved in the common form of acne, acne vulgaris. Excessive holocrine secretion of sebum and keratin triggered by the surge of the steroid hormone testosterone that occurs in both genders at puberty frequently leads to blocked ducts within the gland. Activity of the normal commensal skin bacterium *Propionibacterium acnes* within the blocked duct commonly produces localized inflammation.

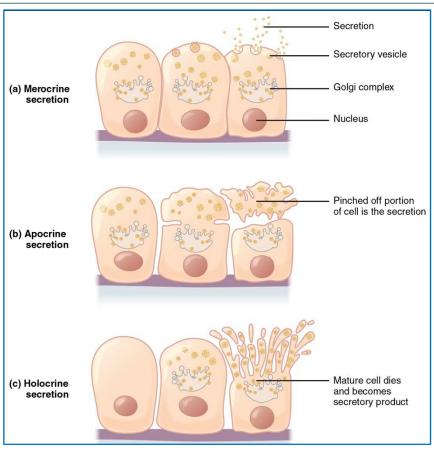


Three basic types of secretion are used by cells of exocrine glands, depending on what substance is being secreted.

- **(a) Merocrine** secretion releases products, usually containing proteins, by means of exocytosis at the apical end of the secretory cells. Most exocrine glands are merocrine.
- **(b) Holocrine** secretion is produced by the disintegration of the secretory cells themselves as they complete their

terminal differentiation, which involves becoming filled with product. Sebaceous glands of hair follicles are the best examples of holocrine glands.

(c) Apocrine secretion involves loss of membrane-enclosed apical cytoplasm, usually containing one or more lipid droplets. Apocrine secretion, along with merocrine secretion, is seen in mammary glands.



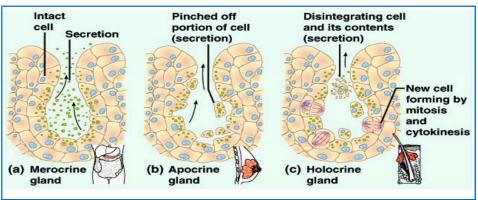
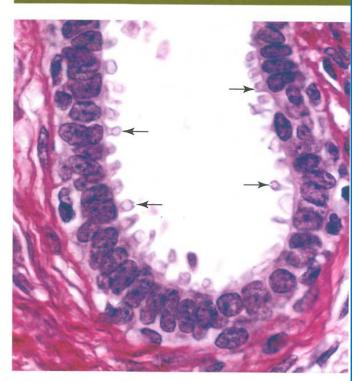


FIGURE **4–22** Holocrine secretion in a sebaceous gland.



In holocrine secretion, best seen in the sebaceous gland adjacent to hair follicles, entire cells fill with a lipid-rich product as they differentiate. Mature (terminally differentiated) cells separate and completely disintegrate, releasing the lipid that serves to protect and lubricate adjacent skin and hair. Sebaceous glands lack myoepithelial cells; cell proliferation inside a dense, inelastic connective tissue capsule continuously forces product into the duct. X200. H&E.

FIGURE **4–23** Apocrine secretion in the mammary gland.



The secretory portions of a mammary gland demonstrate apocrine secretion, characterized by extrusion of the secretion product along with a bit of apical cytoplasm (arrows). The released portion of cell contains lipid droplet(s). Merocrine secretion also occurs from the same and other cells of the gland. X400. PSH.

Merocrine secretion

37

Serous (nonglycosilated Pr.)

Basophilic & acidophilic staining Pancrease & salivary glands

Mucous (glycosilated Pr.)

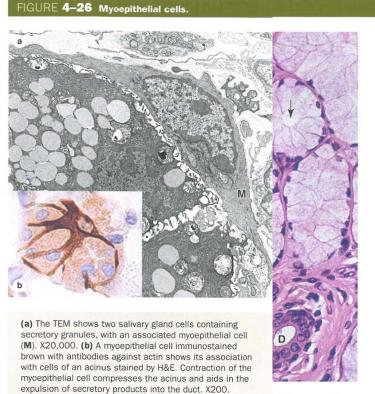
Goblet cells Mucins Eosin PAS

Seromucous glands

Myoephithelial cells

FIGURE 4-24 Serous

The small serous acini of the 5-10 cells facing a very sma is roughly pyramidal, with its light microscopy, the apical the abundant secretory gran ends contain the nuclei and

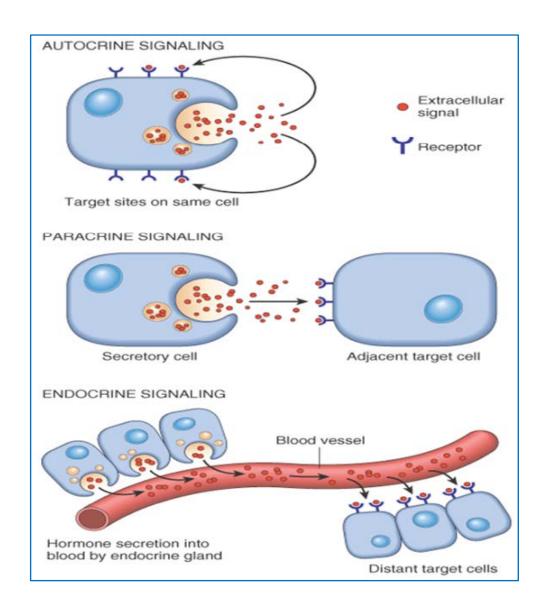


cically larger than serous t of the cytoplasm is ig mucinogen like that inplexes of mucous cells eins with water-binding icous tubules are larger iective tissue surrounds

the mucous tubules and ducts (□). x≥00. PT.

EndocrineProteins (RER)Steroids (SER)

- Paracrine
- Autocrine



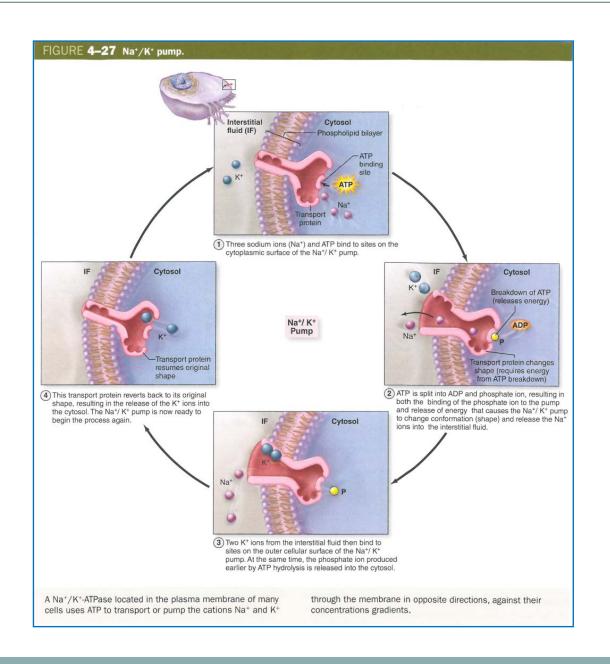
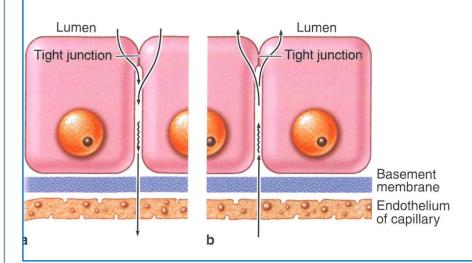


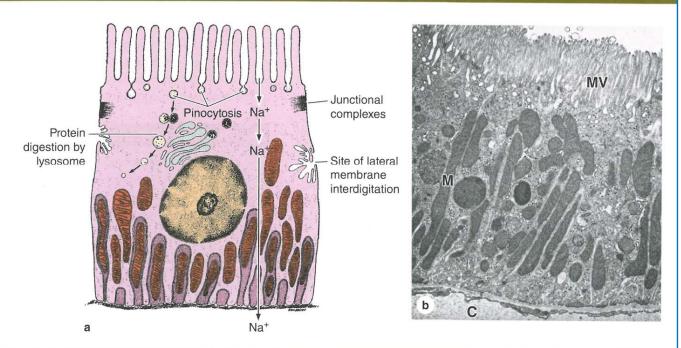
FIGURE 4-28 Ion and water absorption and secretion.



Ion and water transport across epithelia can occur in either direction, depending on which tissue is involved. (a) The direction of transport is from the lumen to the blood vessel, as in the gallbladder and intestine. This process is called **absorption**, and serves to concentrate bile and obtain water and ions in these organs.

(b) Transport of water in the other direction from the capillaries into a lumen, as in the choroid plexus and sweat glands, is often part of **secretion** and serves to expel water from the interstitial fluid into specialized aqueous fluids in these tissues. No matter whether an epithelium is absorbing or secreting water, apical occluding junctions are necessary to maintain tight compartmentalization.

FIGURE 4-29 Absorptive cells.



A diagram and TEM of epithelial cells highly specialized for absorption: cells of proximal convoluted tubule of the kidney. Typically, long invaginations of the basal cell membrane outline regions with mitochondria (M). Interdigitations from neighboring cells are also present laterally. Immediately below the microvilli (MV) are many pinocytotic vesicles, which may fuse with lysosomes as shown or mediate transcytosis by secreting their contents at the basolateral membrane. Junctional

complexes between individual cells separate the apical and basolateral compartments. Sodium ions diffuse passively through the apical membranes of renal epithelial cells and are actively transported out of the cells by Na⁺/K⁺-ATPase located in the basolateral membrane. Immediately below the basal lamina is a capillary (**C**) that removes water absorbed across the epithelium. X9600.

Epithelial cell renewal

42

- Fast (intestine)
- Slow (gland)
- Stem cells in basal part (niche)

>> MEDICAL APPLICATION

Some epithelial cells are prone to abnormal growth or dysplasia, which can progress to precancerous growth called **neoplasia**. Early neoplastic growth is often reversible and does not always result in cancer.

Under certain abnormal conditions, one type of epithelial tissue may undergo transformation into another type in another reversible process called **metaplasia**. In heavy cigarette smokers, the ciliated pseudostratified epithelium lining the bronchi can be transformed into stratified squamous epithelium.

>> MEDICAL APPLICATION

Both benign and malignant tumors can arise from most types of epithelial cells. Malignant tumors of epithelial origin are called **carcinomas** (Gr. *karkinos*, cancer + *oma*, tumor). Malignant tumors derived from glandular epithelial tissue are called **adenocarcinomas** (Gr. *adenos*, gland + *karkinos*). Adenocarcinomas are by far the most common tumors in adults after age 45.

>> MEDICAL APPLICATION

In individuals with chronic vitamin A deficiency, epithelial tissues of the type found in the bronchi and urinary bladder are gradually replaced by stratified squamous epithelium.