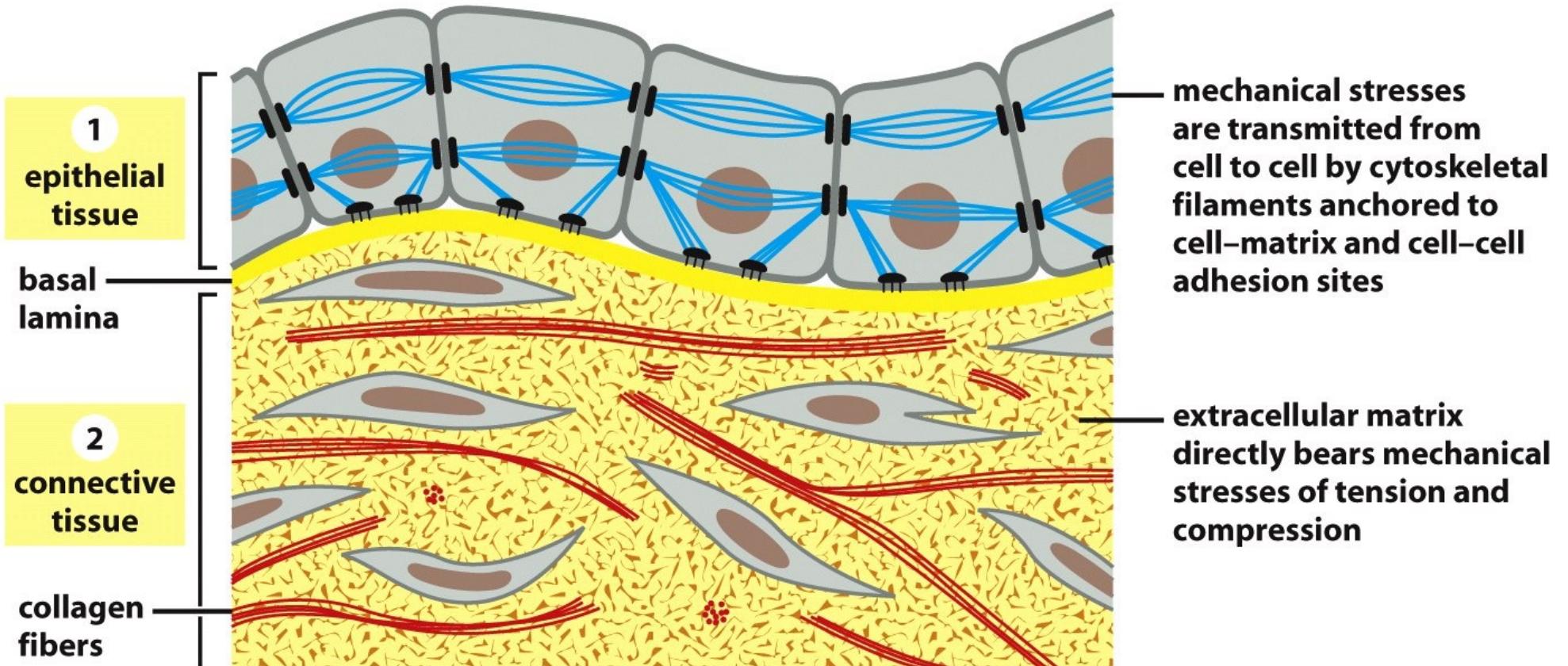


# **Matriz extracelular y uniones celulares**

**Dr. Alejandro Roth**

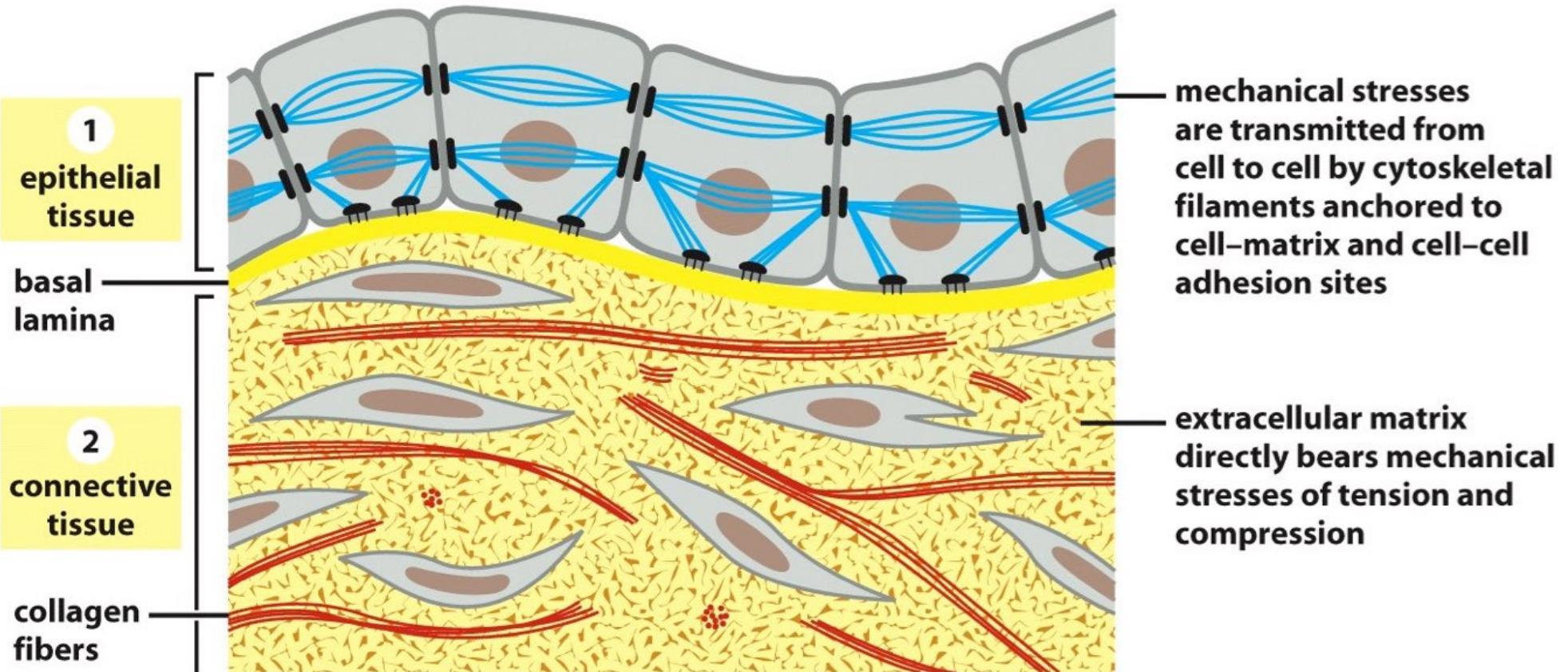
**(Clase basada en la clase del Dr. Christian Gonzalez-Billault)**

# Barreras!



# Organos!

# UNIONES CELULARES



El modelo más utilizado son las células epiteliales debido a que presentan polaridad, sin embargo, hay múltiples casos de gran importancia. Por ejemplo, las células musculares deben poder anclarse a su medio con mucha fuerza.

Lo que vamos a ver....

1.- Matrix extracelular animal y vegetal

2.- Uniones celulares

3.- Resumen

Lo que vamos a ver....

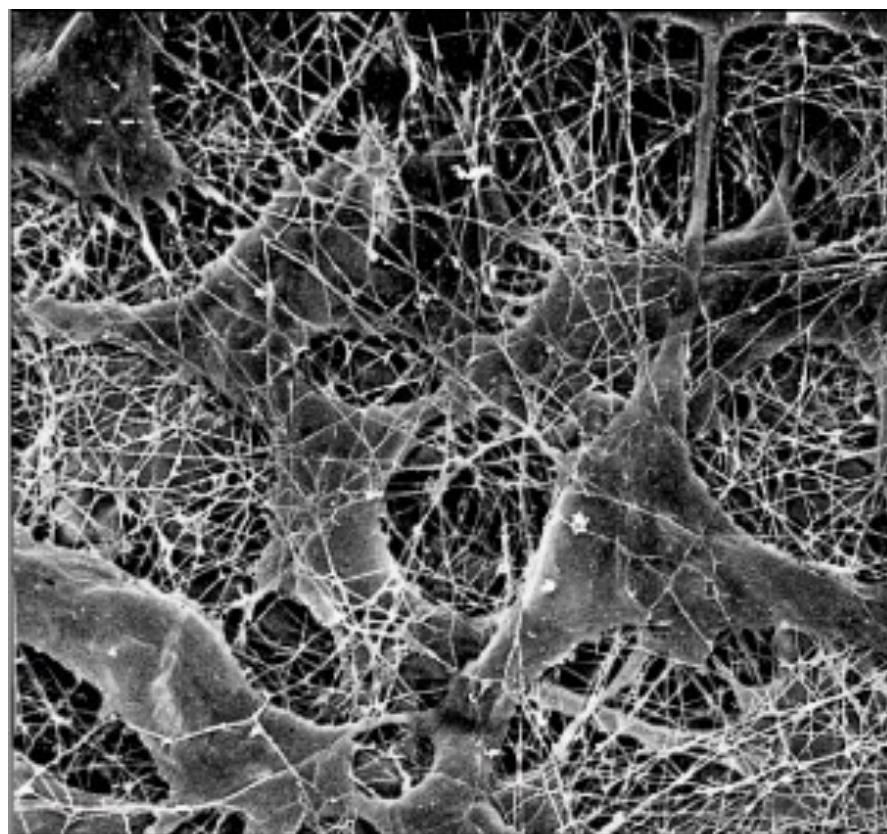
1.- Matrix extracelular animal y vegetal

2.- Uniones celulares

3.- Resumen

## Matrix extracelular?

Cualquier material producido y secretado por las células hacia el medio que las rodea, usualmente se refiere a la porcion no celular de los tejidos



- Celulas secretoras de MEC son los fibroblastos (tejido conectivo)
- En el caso de tejidos conectivos más especializados como cartílago (condroblastos) y hueso (osteoblastos)
  - Esta compuesta de 1) elementos fibrosos (colágeno), 2) proteínas de unión (fibronectina y laminina) y 3) moléculas complejas que llenan el espacio extracelular (proteoglicanos)

## Matrix extracelular?

Cualquier material producido y secretado por las células hacia el medio que las rodea, usualmente se refiere a la porcion no celular de los tejidos

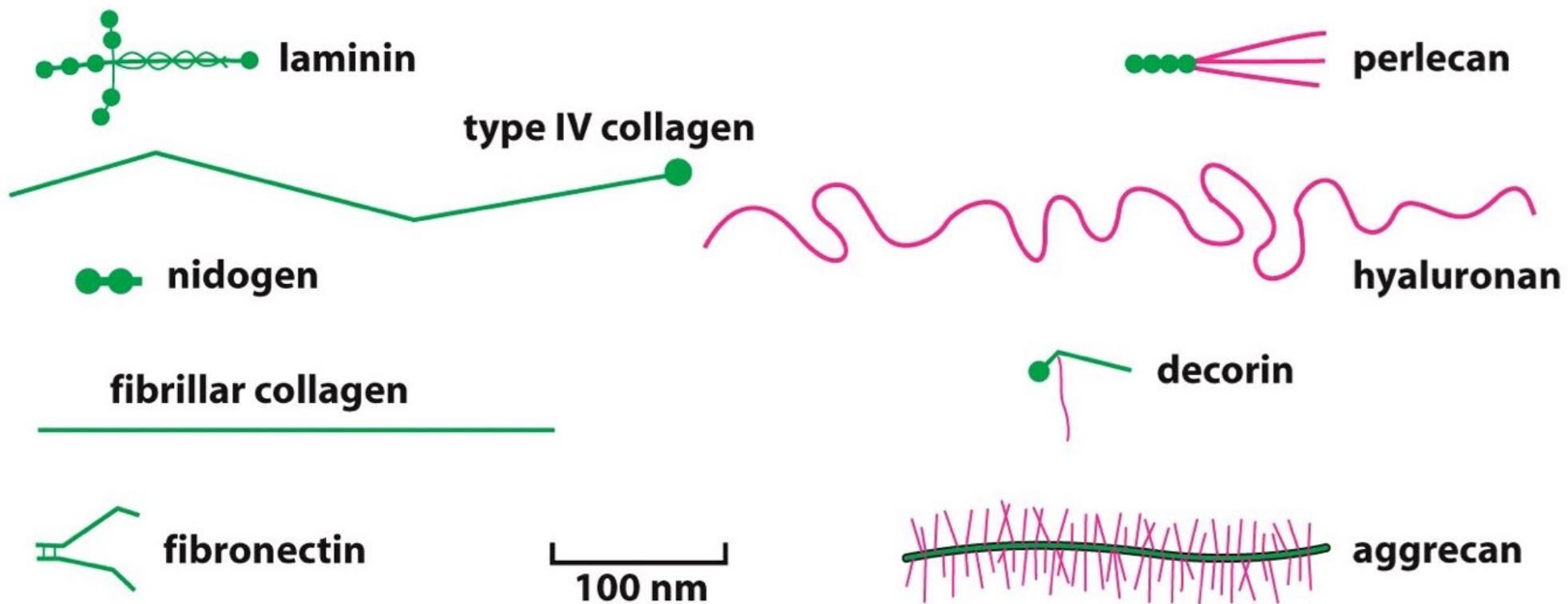


Figure 19-41 Molecular Biology of the Cell 5/e (© Garland Science 2008)

- Esta compuesta de 1) elementos fibrosos (colágeno), 2) proteínas de unión (fibronectina y laminina) y 3) moléculas complejas que rellenan el espacio extracelular (proteoglicanos)

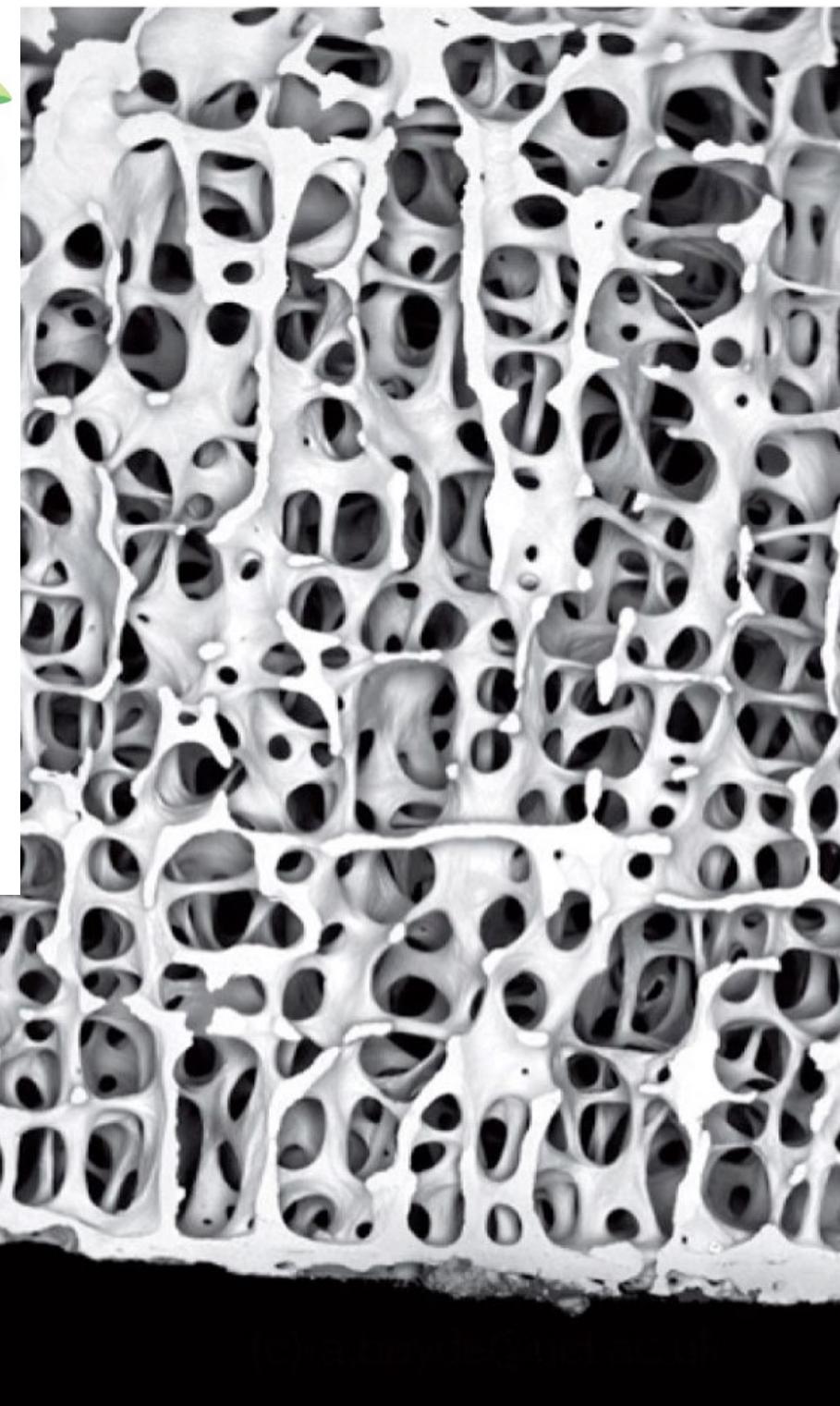
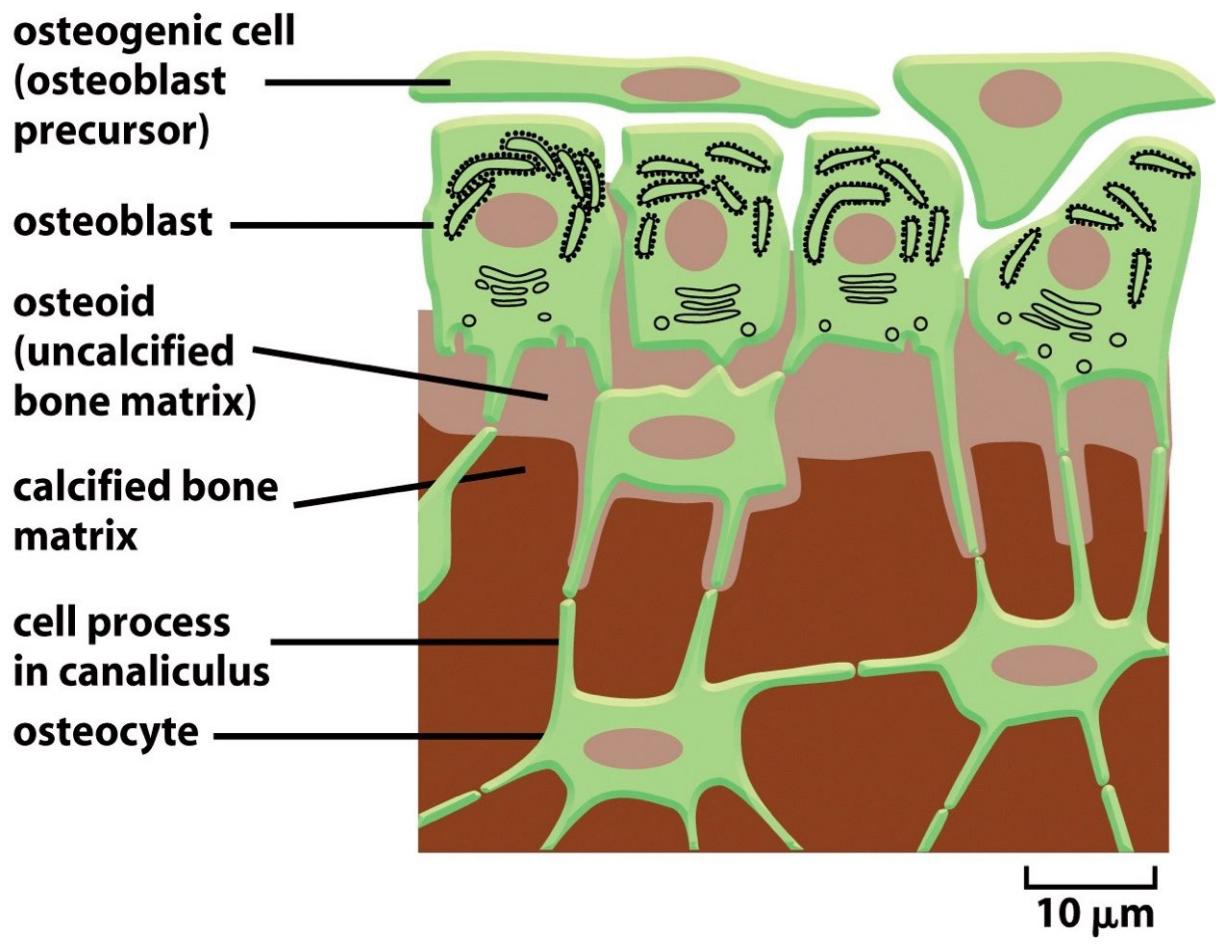


Figure 23-55 *Molecular Biology of the Cell* (© Garland Science 2008)

Figure 23-56a *Molecular Biology of the Cell* (© Garland Science 2008)

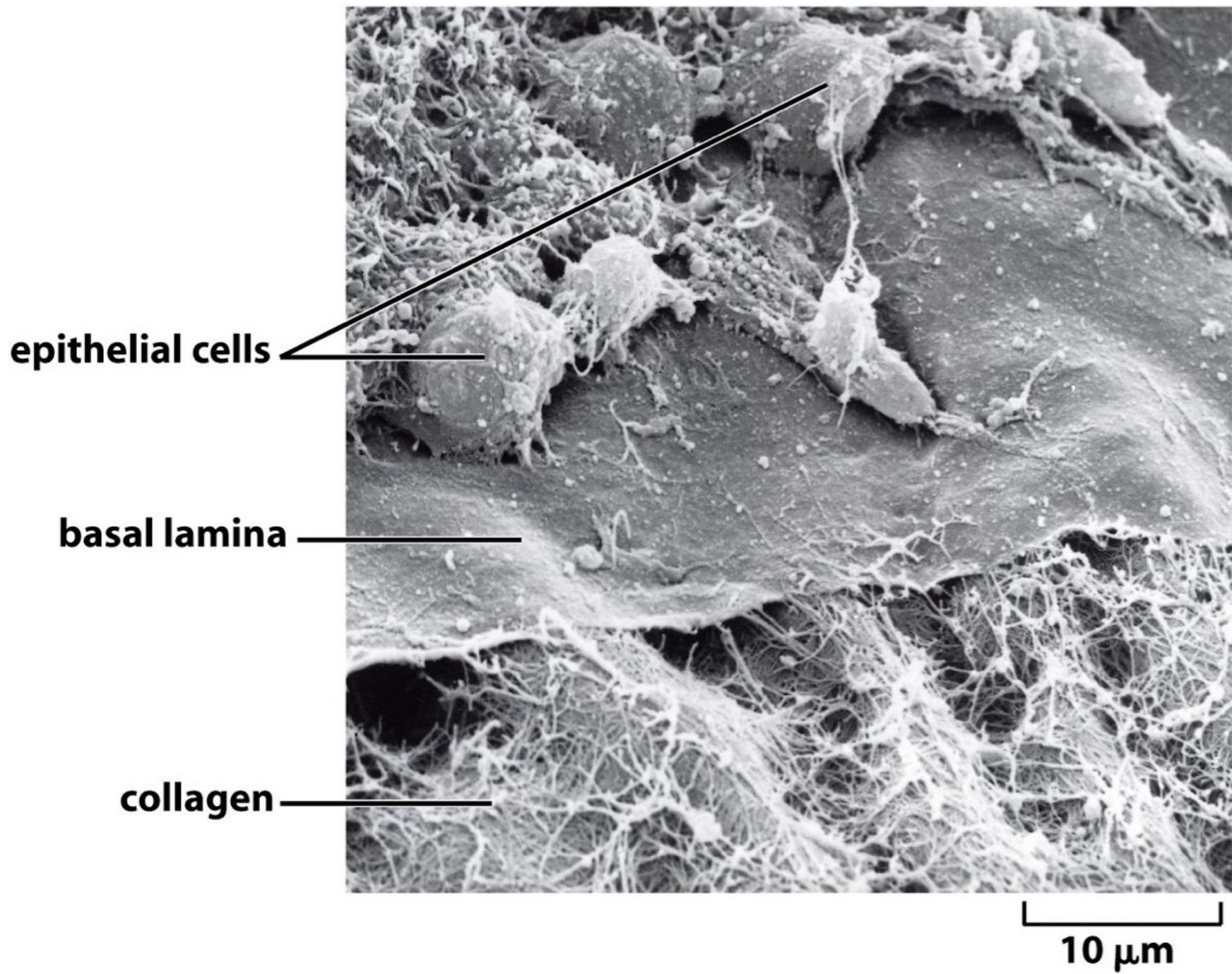


Figure 19-40 Molecular Biology of the Cell 5/e (© Garland Science 2008)

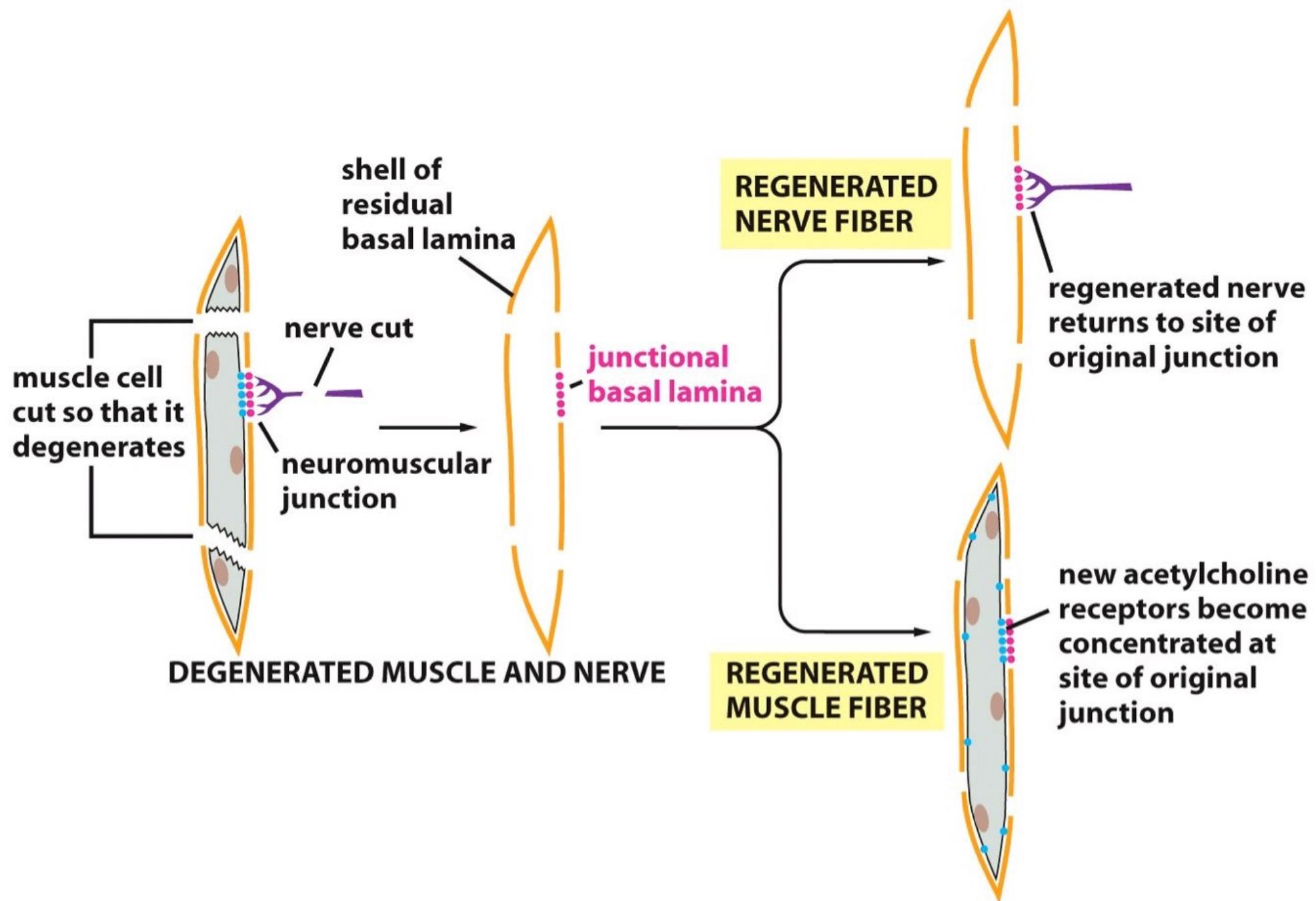
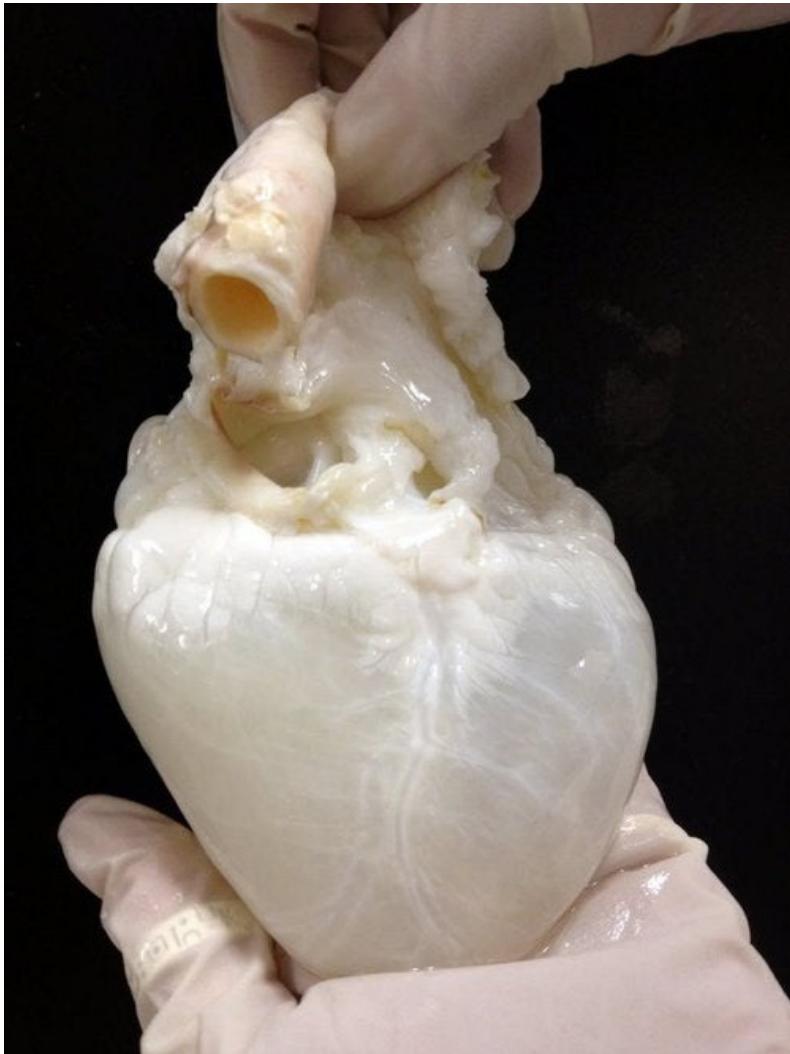


Figure 19-44 Molecular Biology of the Cell 5/e (© Garland Science 2008)



## Corazón “Fantasma”

Corazón humano de-celularizado. Solo queda la matriz extracelular que será repoblada con células troncales del receptor. Estas se diferenciarían a músculo cardíaco o células vasculares.

Se disminuye la posibilidad de rechazo del tejido, por lo que aumentan los órganos disponibles.

<https://i.imgur.com/yb7BH0A.jpg>

### Tissue Engineering: How to Build a Heart

With thousands of people in need of heart transplants, researchers are trying to grow new organs

By Brendan Maher, Nature magazine on July 8, 2013

### Decellularized matrices in regenerative medicine

Doris A. Taylor, Luiz C. Sampaio, Zannatul Ferdous, Andrea S. Gobin, Lakeshia J. Taitea

<https://doi.org/10.1016/j.actbio.2018.04.044>

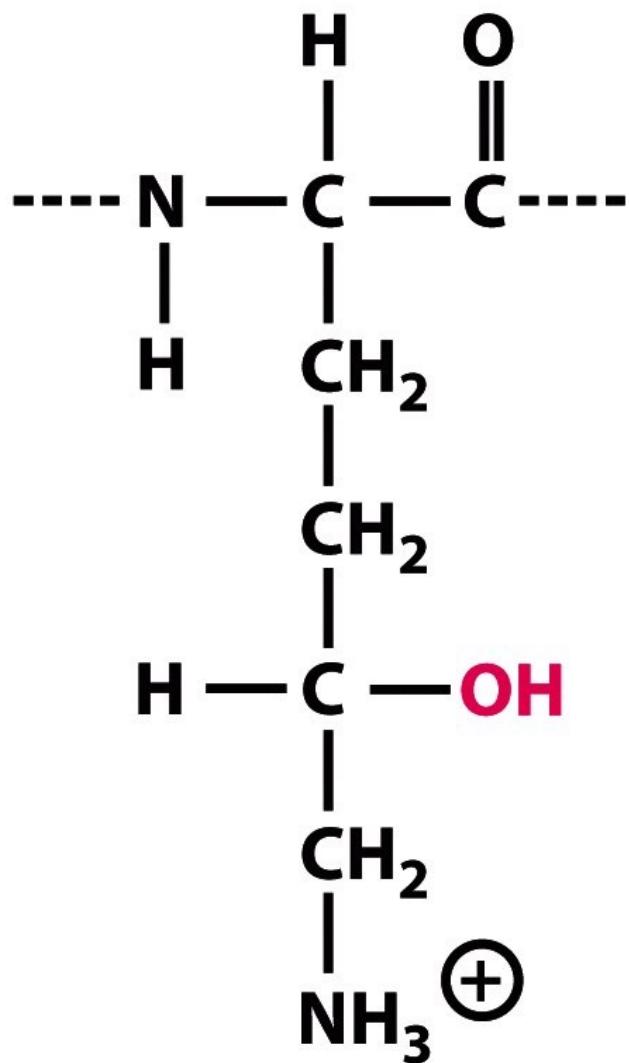
# 1.- Elementos fibrosos (colageno)



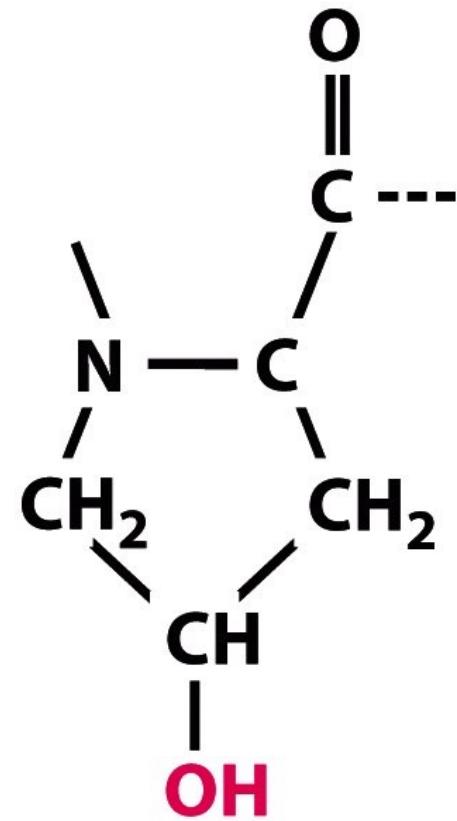
**Table 19–7 Some Types of Collagen and Their Properties**

	TYPE	POLYMERIZED FORM	TISSUE DISTRIBUTION	MUTANT PHENOTYPE
Fibril-forming (fibrillar)	I	fibril	bone, skin, tendons, ligaments, cornea, internal organs (accounts for 90% of body collagen)	severe bone defects, fractures
	II	fibril	cartilage, invertebral disc, notochord,	cartilage deficiency, dwarfism vitreous humor of the eye
	III	fibril	skin, blood vessels, internal organs	fragile skin, loose joints, blood vessels prone to rupture
	V	fibril (with type I)	as for type I	fragile skin, loose joints, blood vessels prone to rupture
	XI	fibril (with type II)	as for type II	myopia, blindness
Fibril-associated	IX	lateral association	cartilage with type II fibrils	osteoarthritis
Network-forming	IV	sheetlike network	basal lamina	kidney disease (glomerulonephritis), deafness
Transmembrane	VII	anchoring fibrils	beneath stratified squamous epithelia	skin blistering
	XVII	non-fibrillar	hemidesmosomes	skin blistering
Proteoglycan core protein	XVIII	non-fibrillar	basal lamina	myopia, detached retina, hydrocephalus

Note that types I, IV, V, IX, and XI are each composed of two or three types of  $\alpha$  chains (distinct, nonoverlapping sets in each case), whereas types II, III, VII, XII, XVII, and XVIII are composed of only one type of  $\alpha$  chain each. Only 10 types of collagen are shown, but about 27 types of collagen and 42 types of  $\alpha$  chains have been identified in humans.



**hydroxylysine  
in protein**



**hydroxyproline  
in protein**

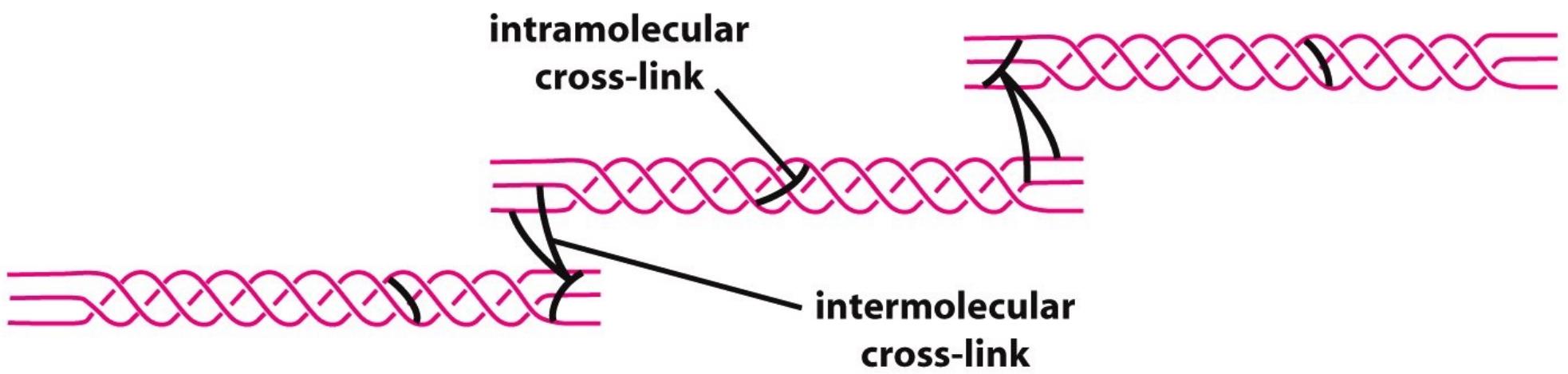


Figure 19-65 *Molecular Biology of the Cell* (© Garland Science 2008)

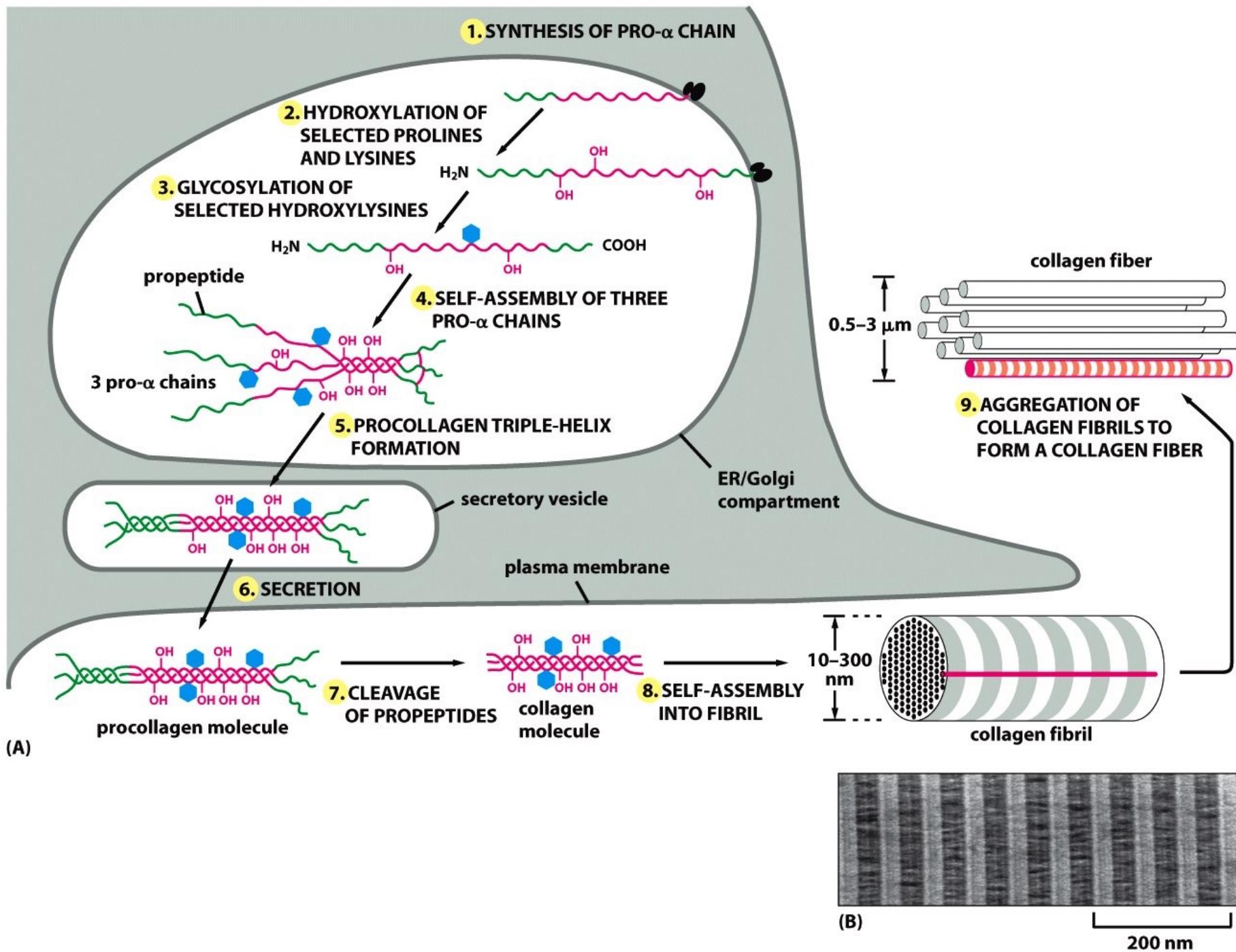
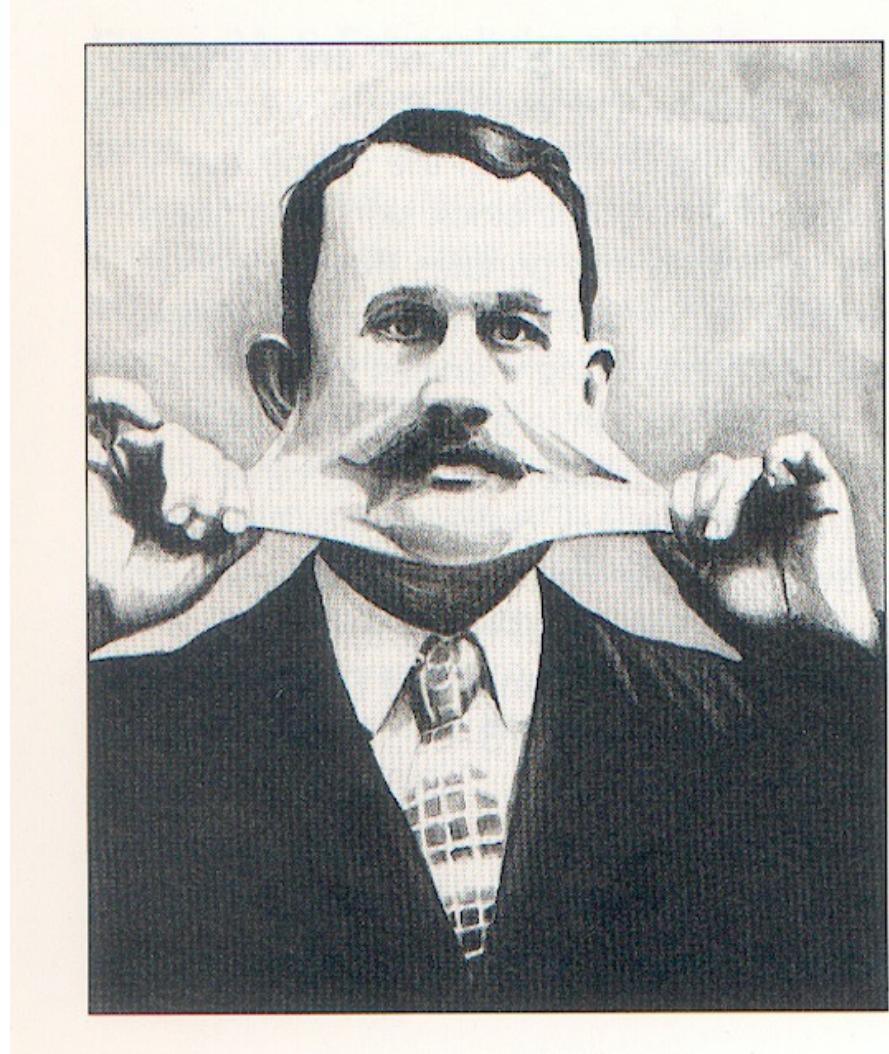
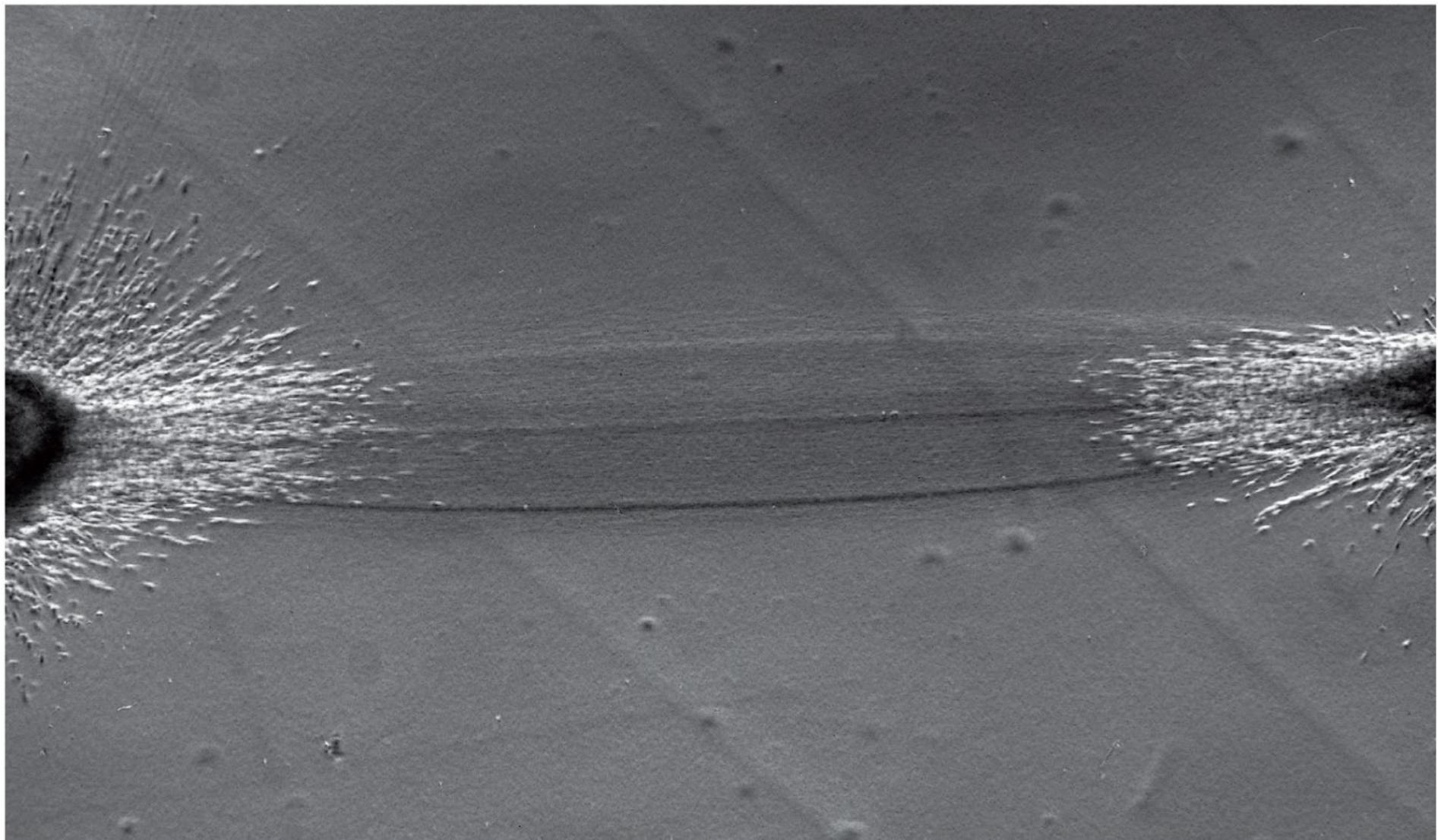


Figure 19-66 Molecular Biology of the Cell (© Garland Science 2008)



James Morris (1890)

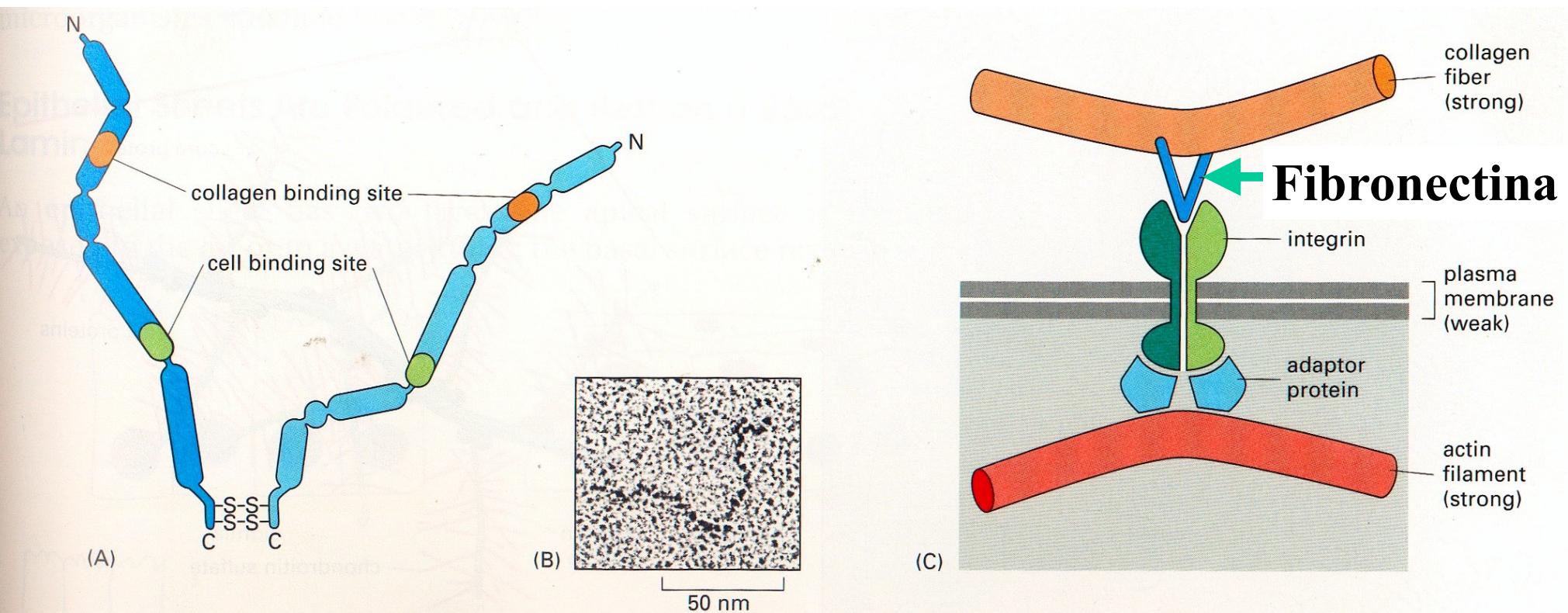


A horizontal scale bar consisting of two thick black L-shaped brackets with a gap in the center, representing a length of one millimeter.

**1 mm**

Figure 19-69 *Molecular Biology of the Cell* (© Garland Science 2008)

## 2) proteínas de unión (fibronectina y laminina)



**Figure 21-14 Integrins link the extracellular matrix to the cytoskeleton in an animal cell.** (A) Diagram and (B) electron micrograph of a fibronectin molecule. (C) The transmembrane linkage mediated by an integrin molecule. The integrin molecule transmits tension across the plasma membrane: it is anchored inside the cell to the cytoskeleton and externally via fibronectin to the extracellular matrix. The plasma membrane itself does not have to be strong. The integrin shown links fibronectin to an actin filament inside the cell, but other integrins connect different extracellular proteins to the cytoskeleton (usually to actin filaments, but sometimes to intermediate filaments). (B, from J. Engel et al., *J. Mol. Biol.* 150:97–120, 1981. © Academic Press.)

# Laminina

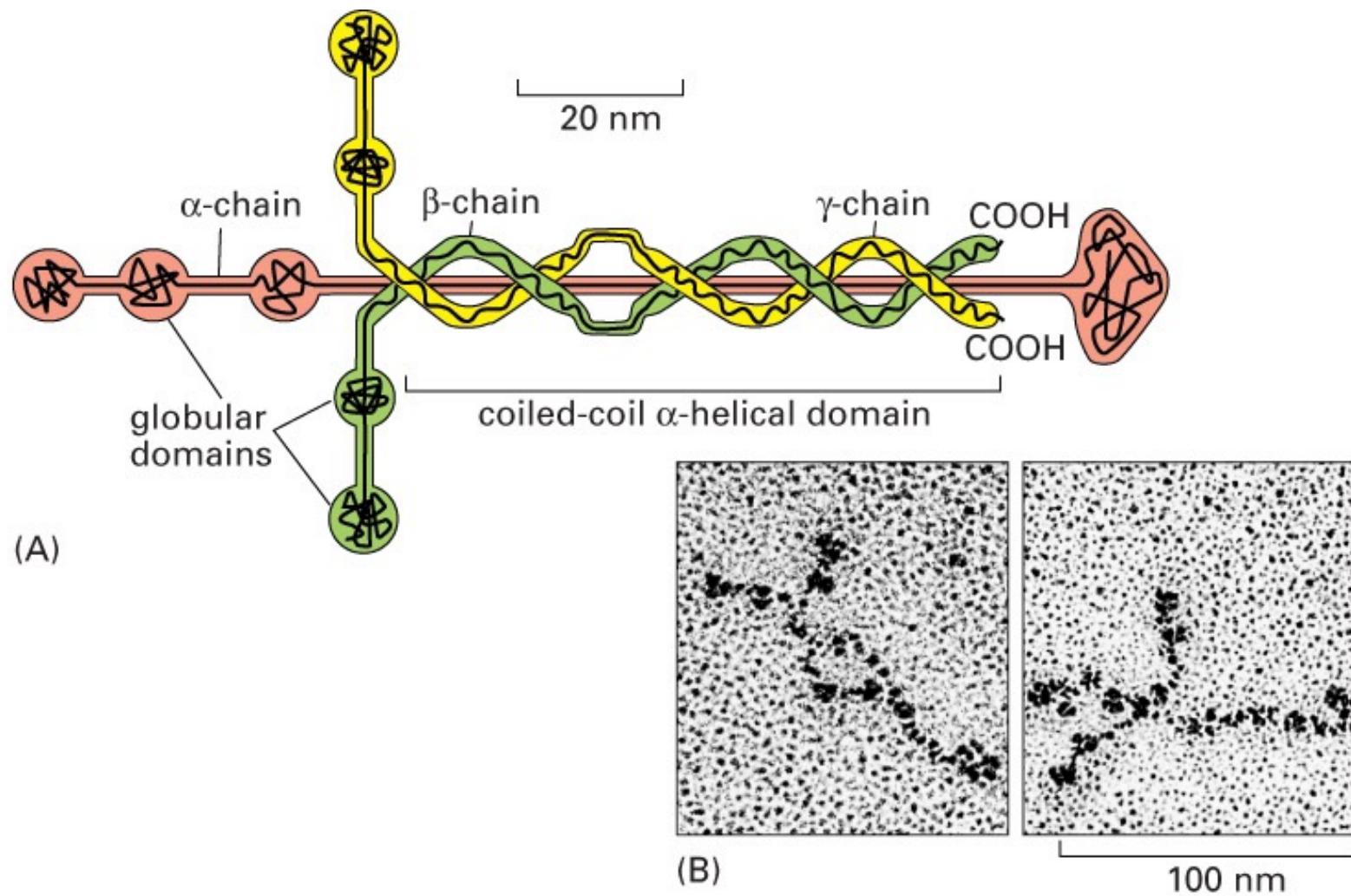
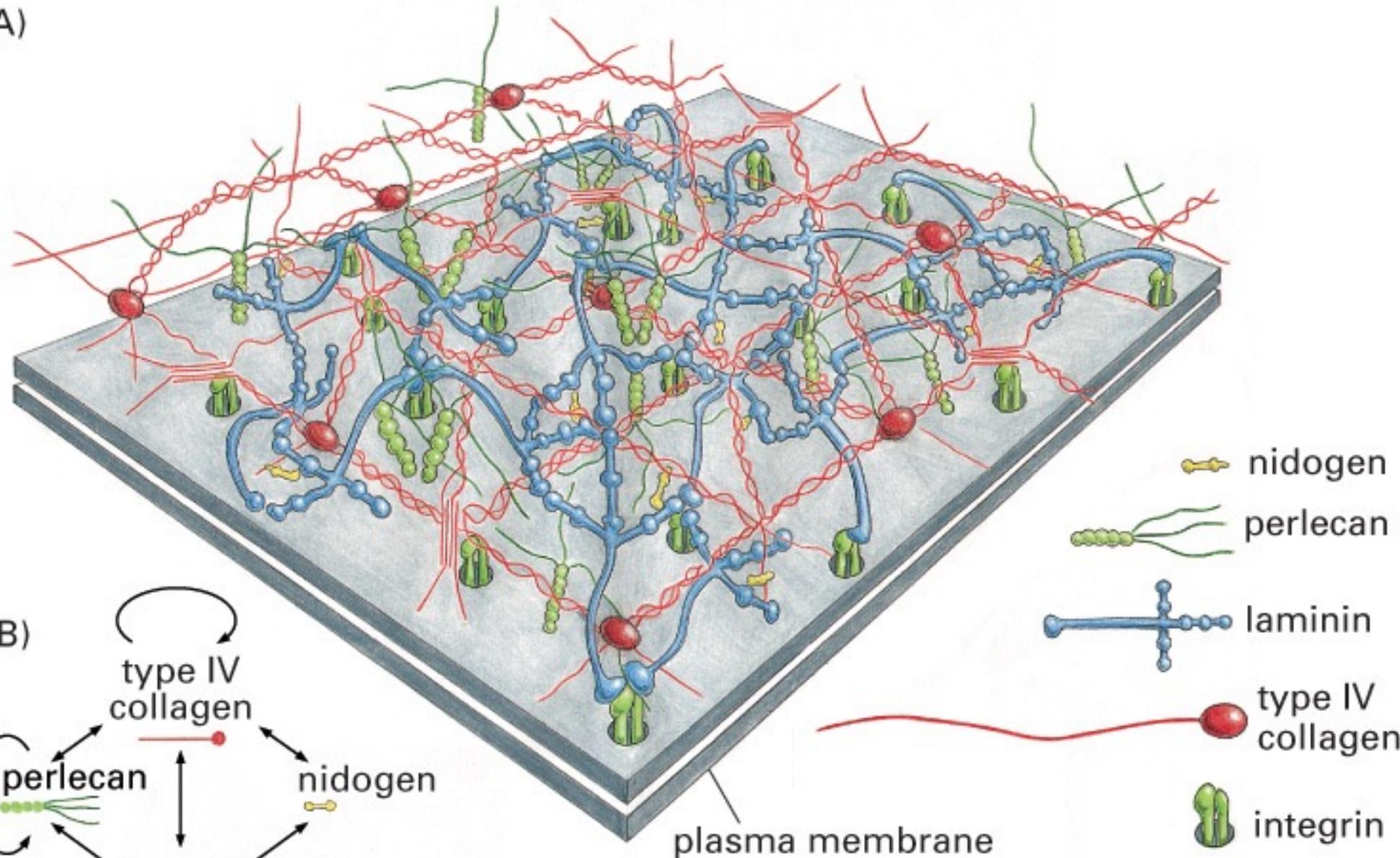


Figure 19–57. Molecular Biology of the Cell, 4th Edition.

(A)



(B)

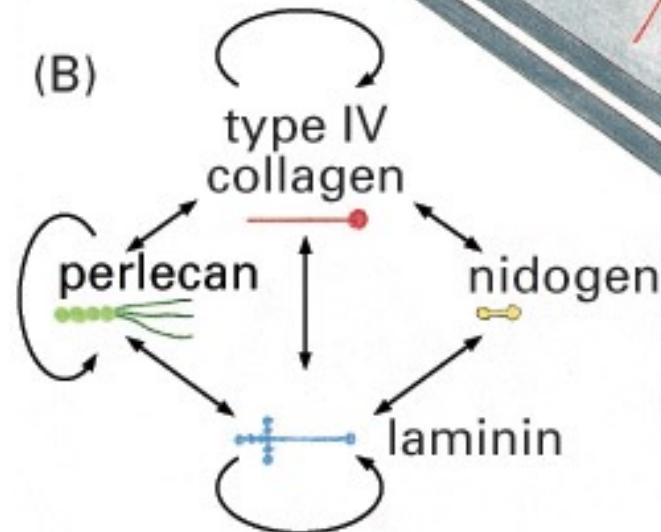


Figure 19–58. Molecular Biology of the Cell, 4th Edition.

### 3) moléculas complejas que llenan el espacio extracelular

globular protein (MW 50,000)

Proteo-glicanos

perlecan

decorin

aggregan

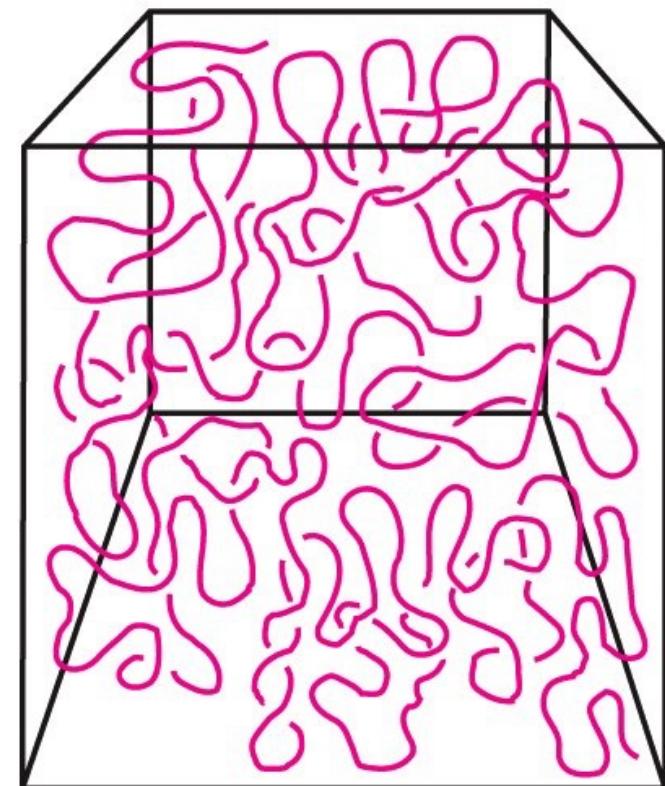


glycogen (MW ~ 400,000)



spectrin (MW 460,000)

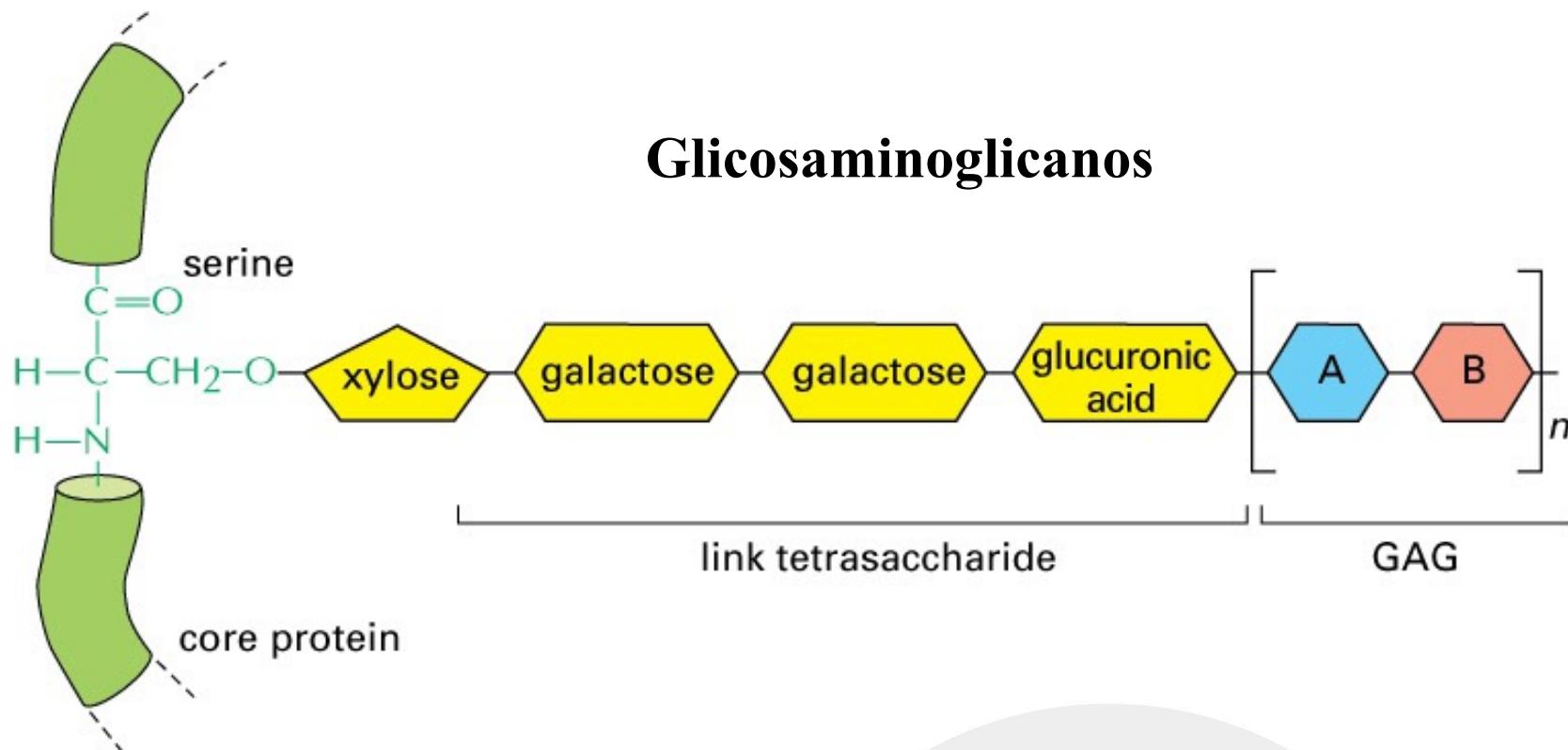
collagen (MW 290,000)



hyaluronan (MW  $8 \times 10^6$ )

300 nm

### 3) moléculas complejas que rellenan el espacio extracelular



**BIO-LIFTER**

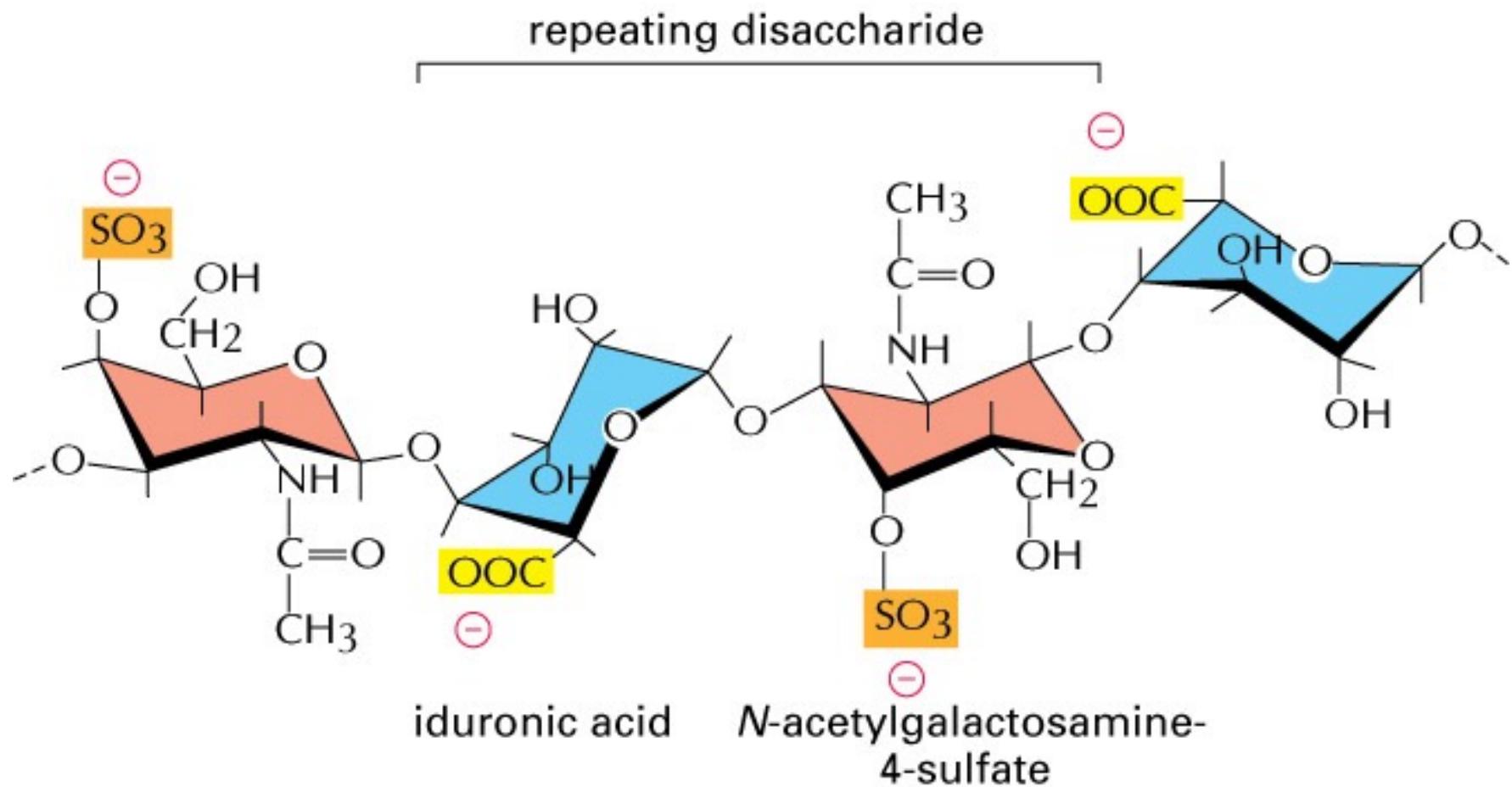
ANTIAGE. HIDRATANTE  
30ml



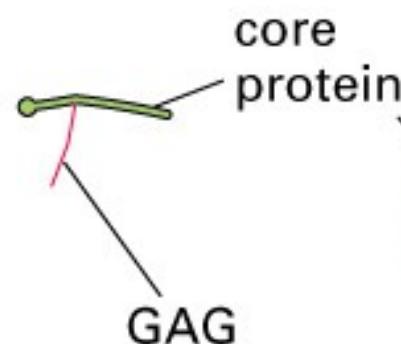
**PRECURSORES DE GAGs, FITOECDISTEROIDES,  
DERIVADOS TRITERPÉNICOS, CERAMIDAS,  
ÁCIDO HIALURÓNICO, SILICONAS REFRACTARIAS.**

- Rejuvenece la arquitectura cutánea.
- Estimula la síntesis de GAGs, proteoglicanos, colágeno y laminina.
- Fortalece la unión dermo-epidérmica.
- Con Ceramidas, Ácido Hialurónico y siliconas refractarias que disimulan imperfecciones.
- Apto para todo tipo de piel.

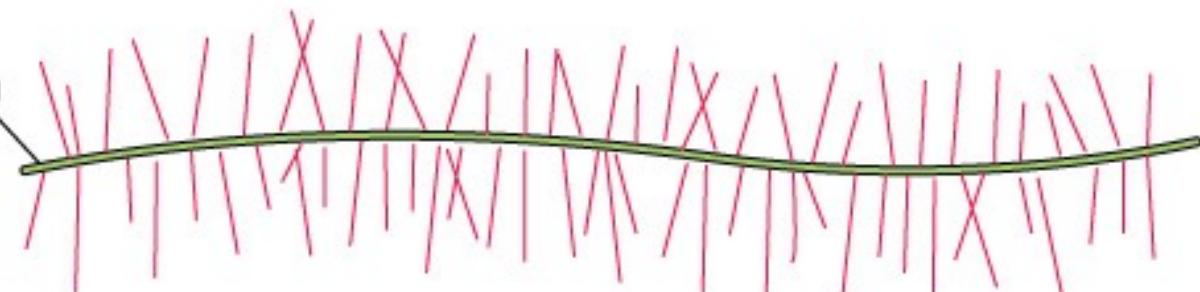
Aplicar directo en el rostro y contorno de ojos de mañana



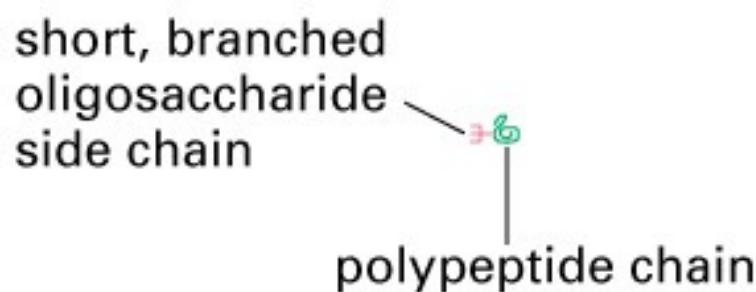
**DECORIN**  
(MW ~ 40,000)



**AGGREGCAN**  
(MW ~  $3 \times 10^6$ )

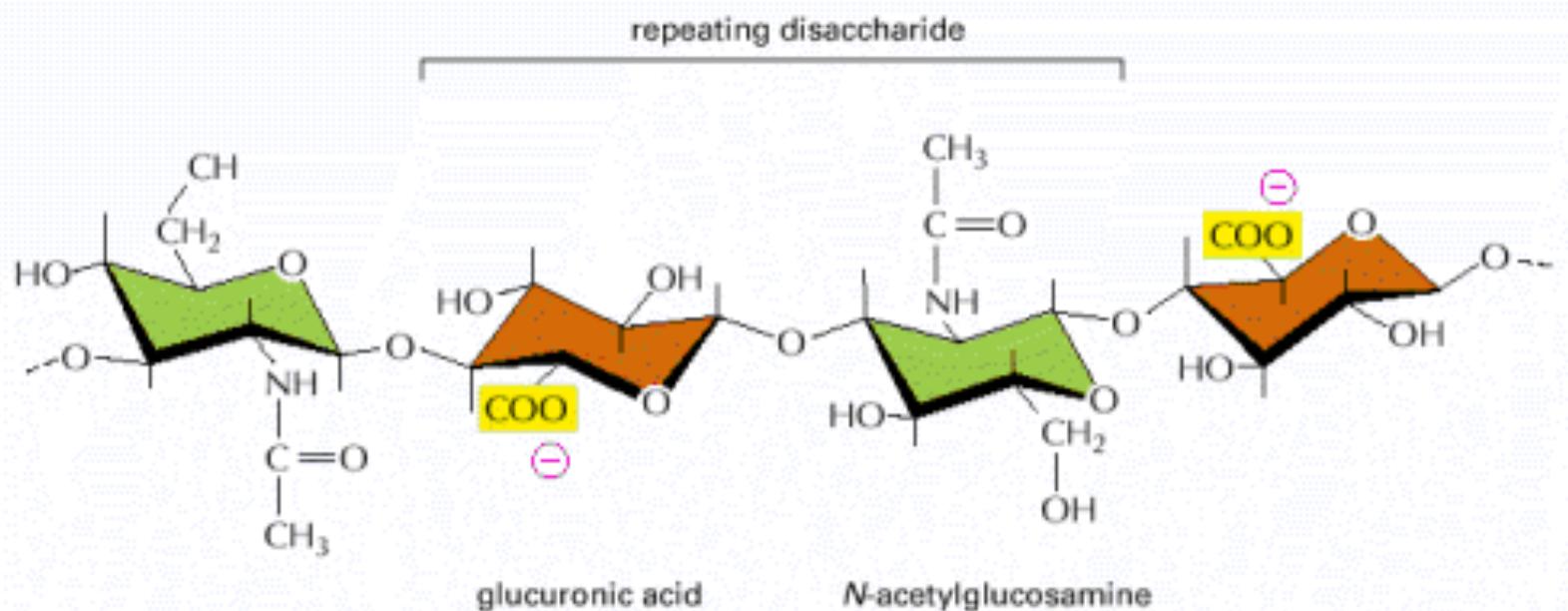


**RIBONUCLEASE**  
(MW ~ 15,000)

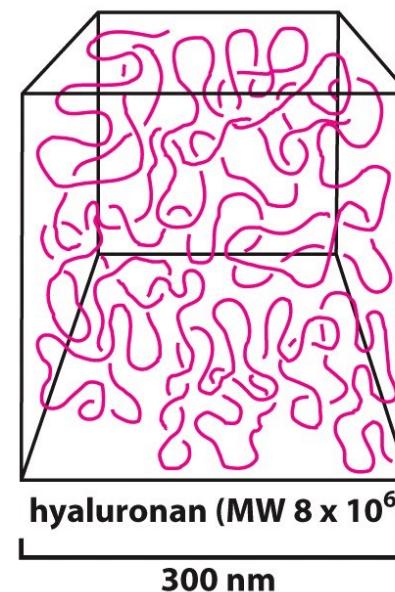


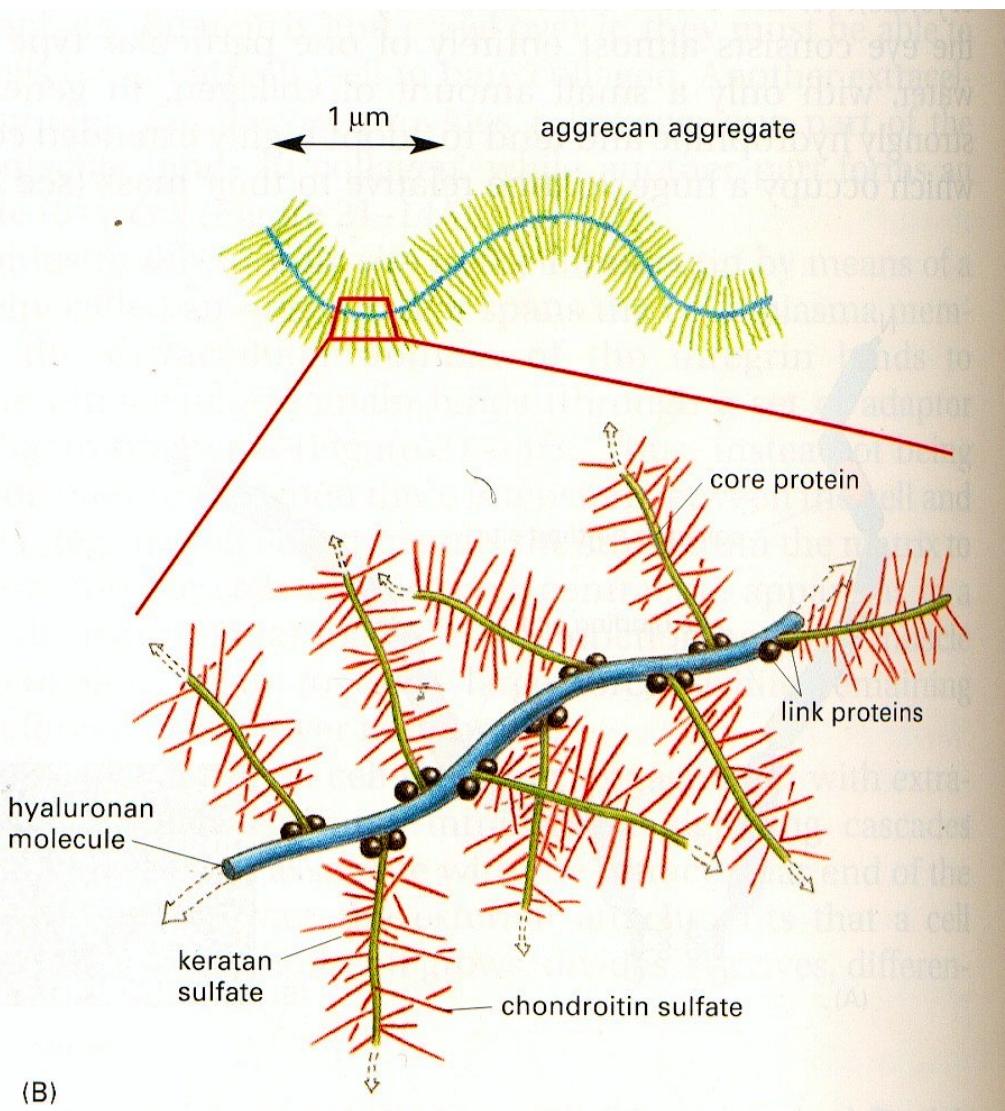
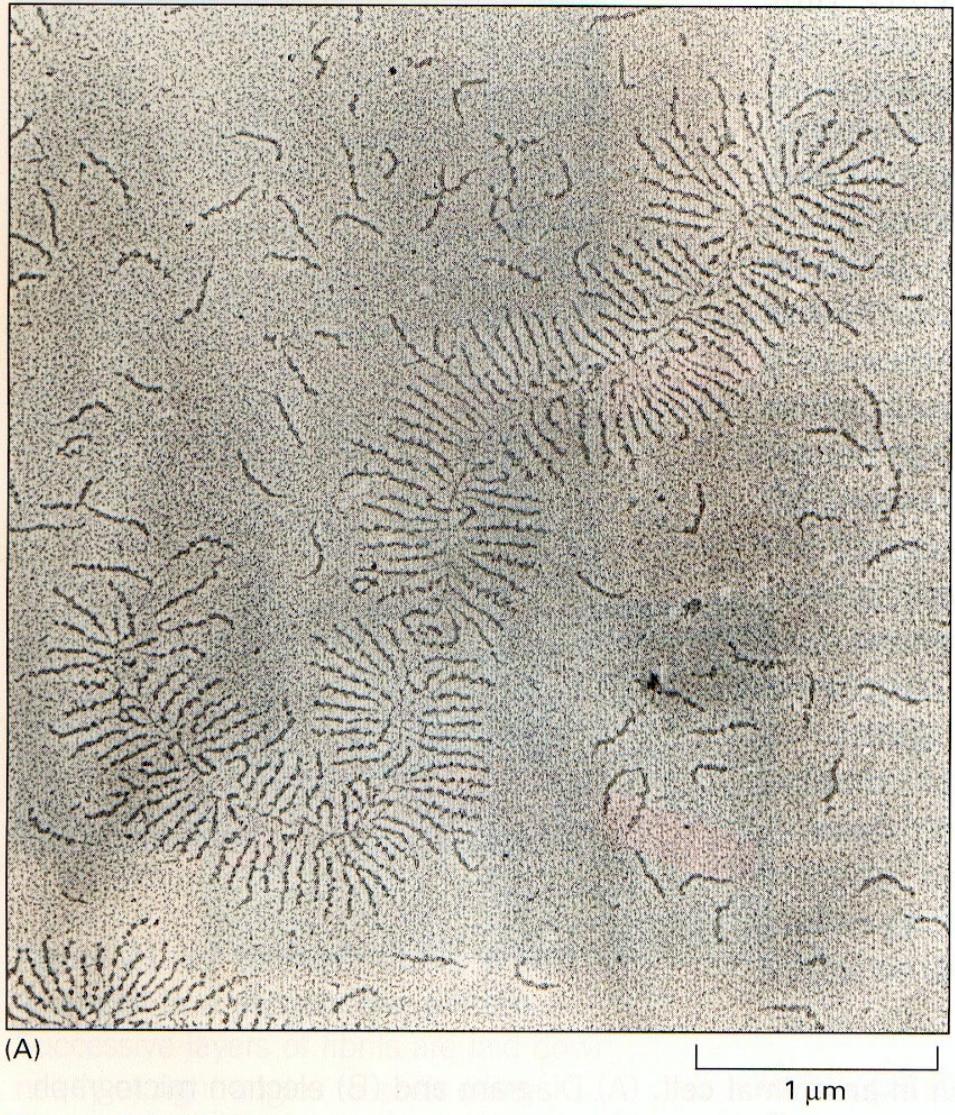
100 nm

# Hialuronano (Ac. Hialouronico)



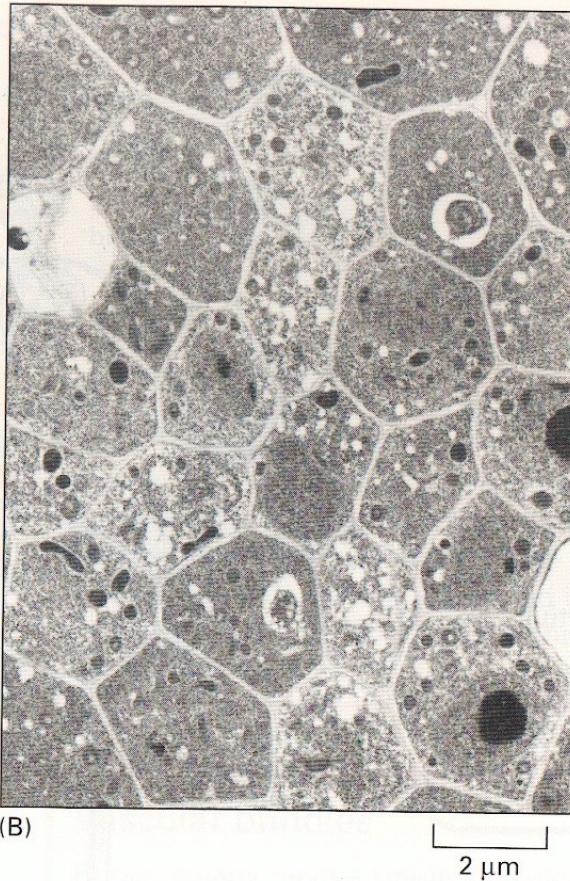
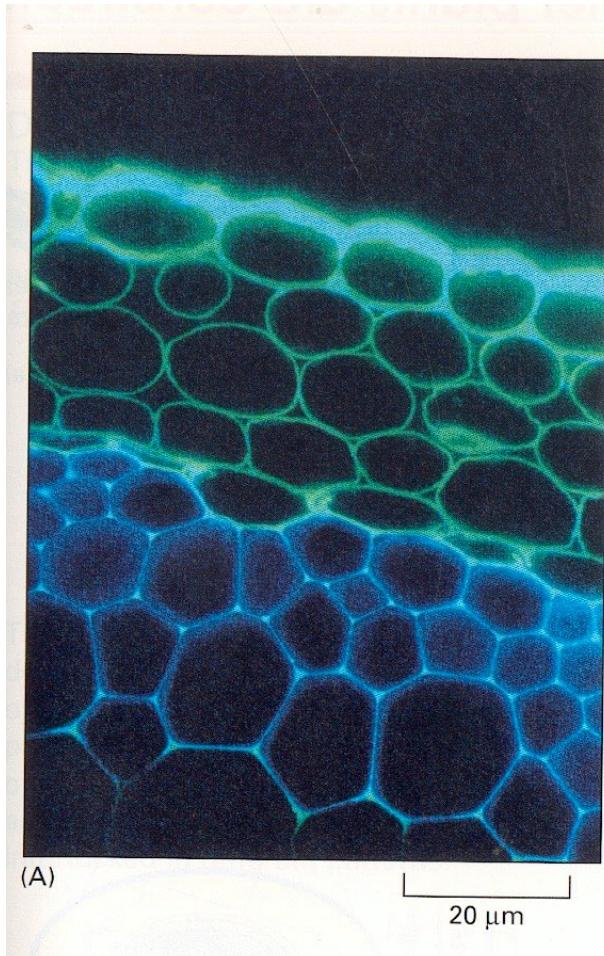
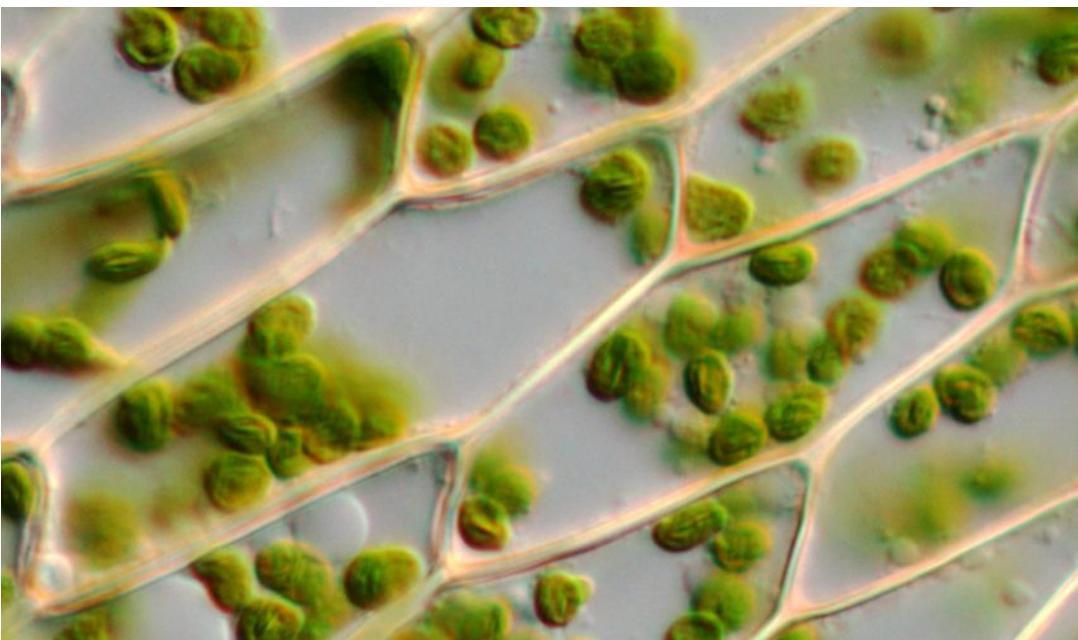
- Mas simple
- No sulfato
- No unido a proteínas





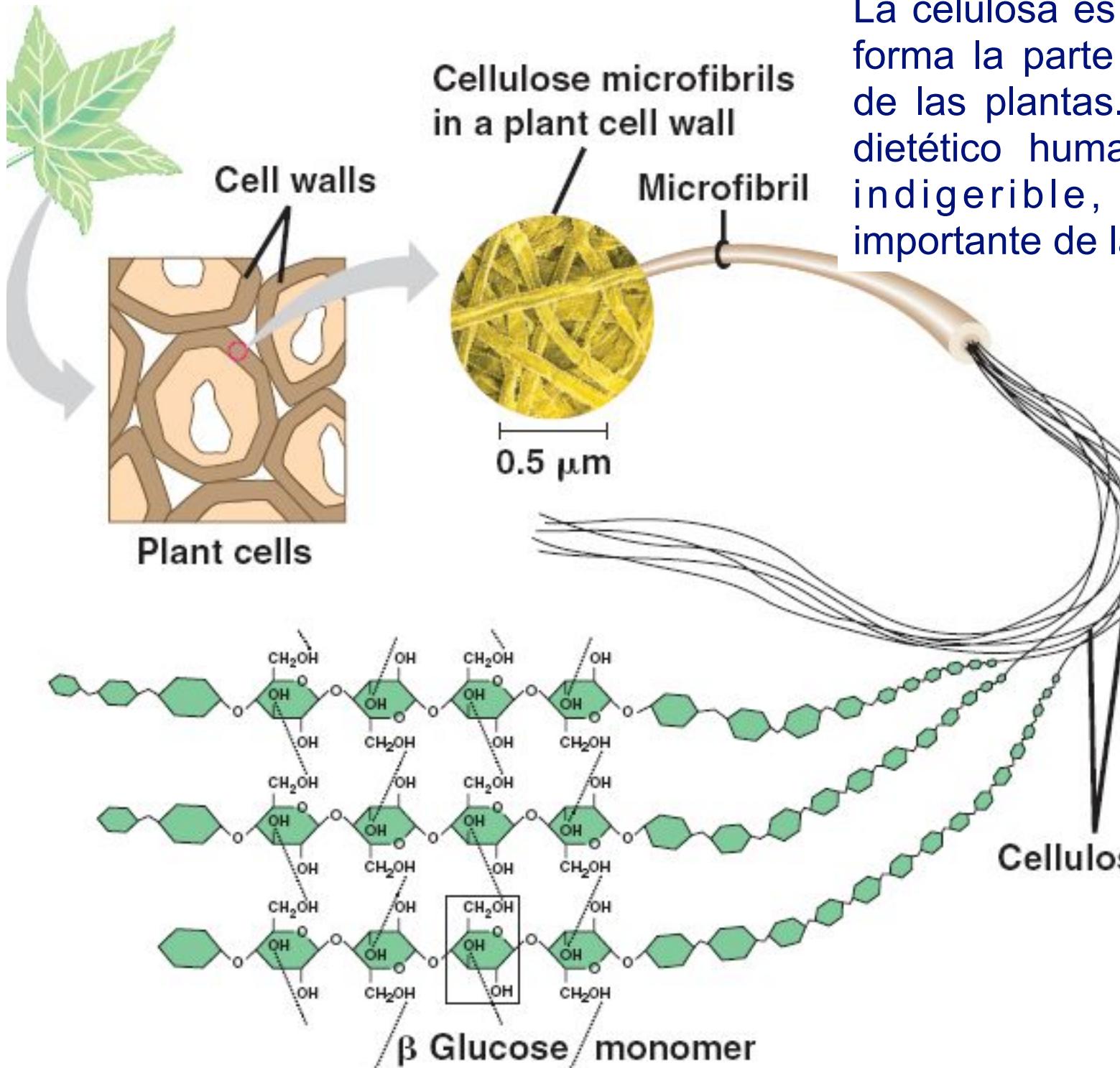
**Figure 21–16 Proteoglycans and GAGs can form large aggregates.** (A) Electron micrograph of an aggregate from cartilage spread out on a flat surface. Many free subunits—themselves large proteoglycan molecules—are also seen. (B) Schematic drawing of the giant aggregate illustrated in (A), showing how it is built up from GAGs and proteins. The molecular weight of such a complex can be  $10^8$  daltons or more, and it occupies a volume equivalent to that of a bacterium, which is about  $2 \times 10^{-12} \text{ cm}^3$ . (A, courtesy of Lawrence Rosenberg.)

# Matrix extracelular vegetal



**Figure 21–3** Plant tissues are strengthened by the plant cell wall.  
(A) A cross section of part of the stem of the flowering plant *Arabidopsis* is shown, stained with fluorescent dyes that label two different cell wall components—cellulose in blue, and another polysaccharide (pectin) in green. The cells themselves are unstained and invisible in this preparation. Regions rich in both cellulose and pectin appear white. Pectin predominates in the outer layers of cells, which have only primary cell walls (deposited while the cell is still growing). Cellulose is more plentiful in the inner layers, which have thicker, more rigid secondary cell walls (deposited after cell growth has ceased). (B) The cells and their walls are clearly seen in this electron micrograph of young cells in the root of the same plant. (Courtesy of Paul Linstead.)

# POLISACÁRIDOS SIMPLES



## CELULOSA

La celulosa es un polisacárido que forma la parte fibrosa de la pared de las plantas. Del punto de vista dietético humano, la celulosa es indigerible, formando parte importante de la fibra dietaria.

La fibra dietaria es un potente anti-cancerígeno en el intestino.

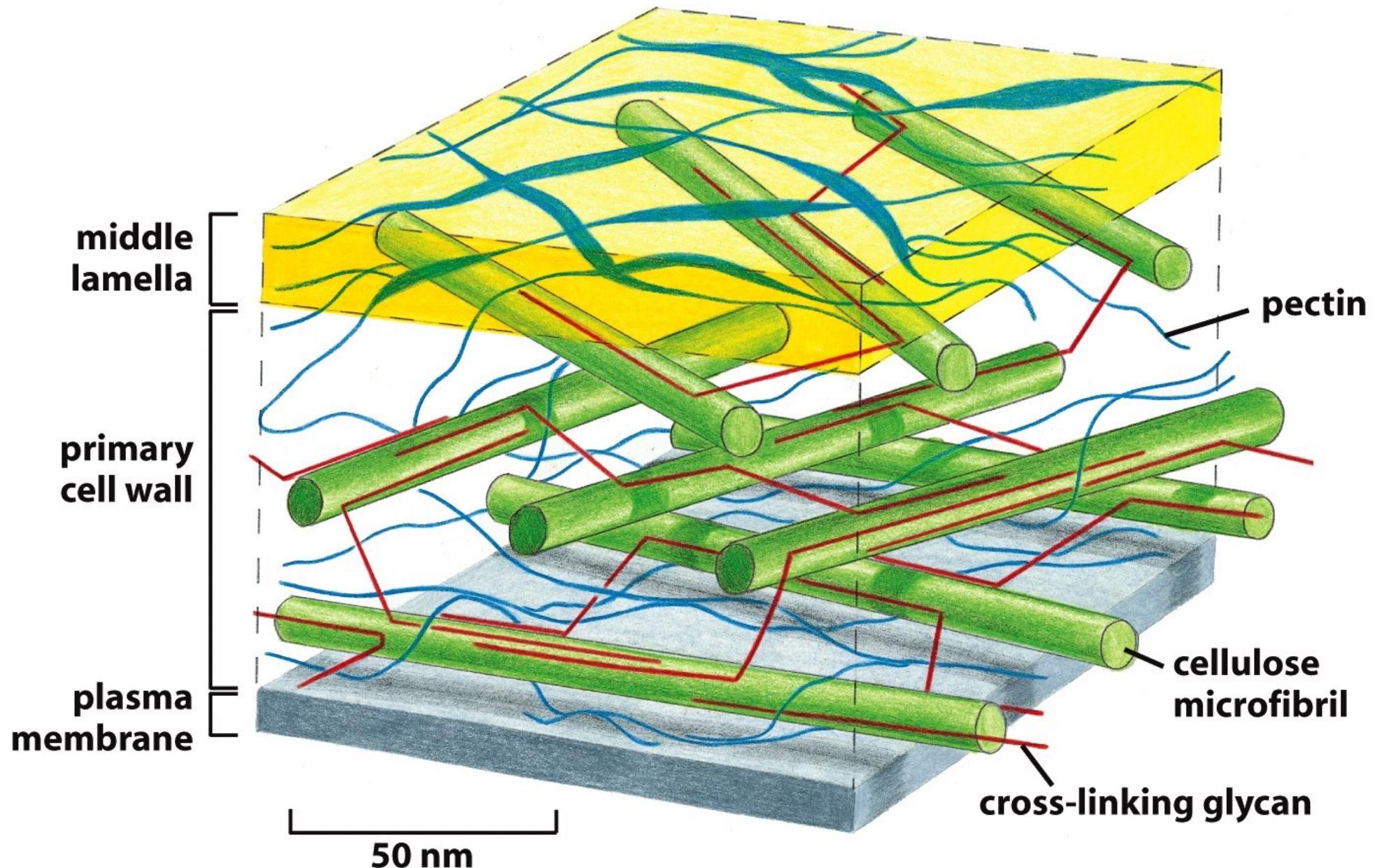
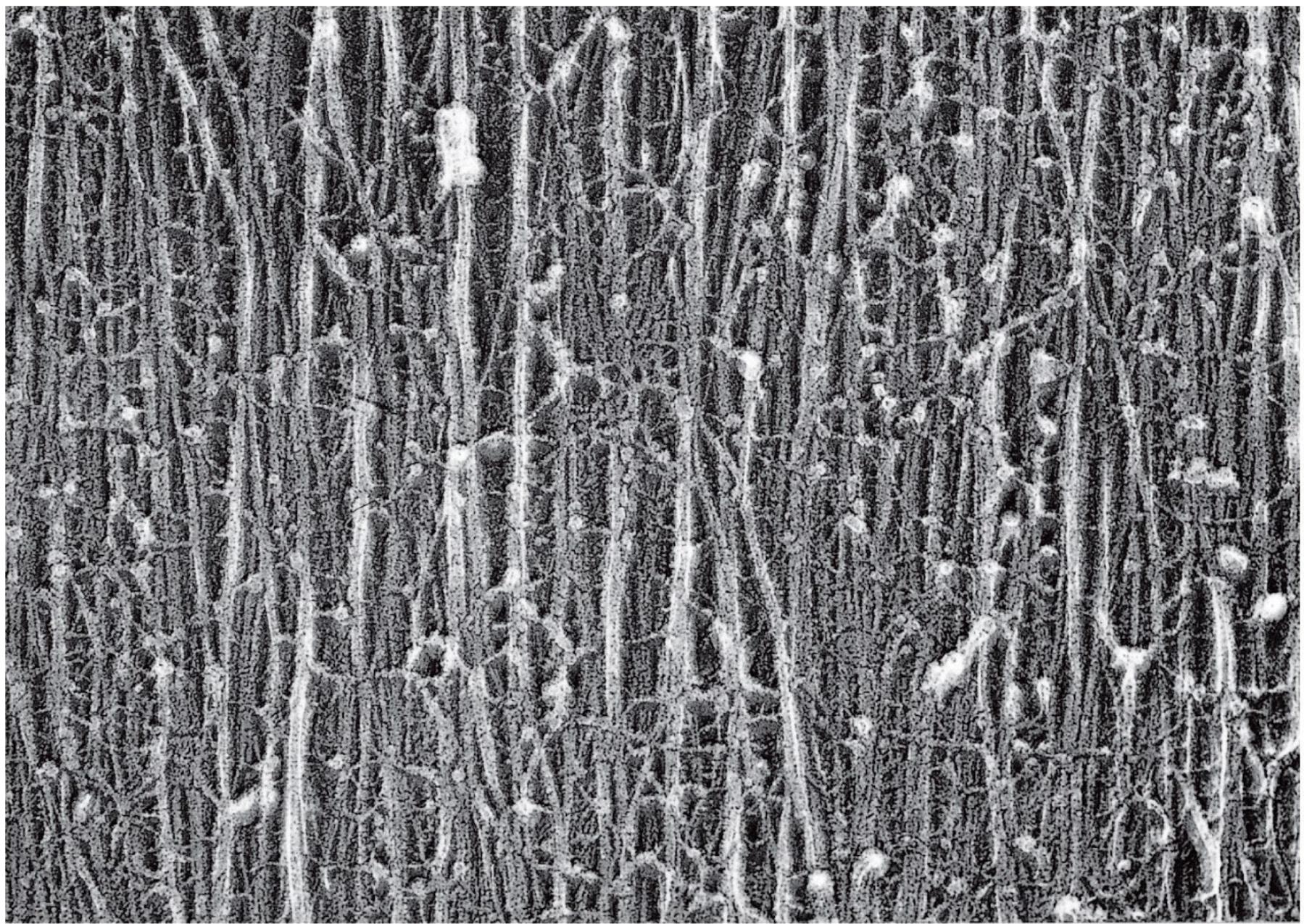


Figure 19-79 *Molecular Biology of the Cell* (© Garland Science 2008)



200 nm

Figure 19-80a *Molecular Biology of the Cell* (© Garland Science 2008)

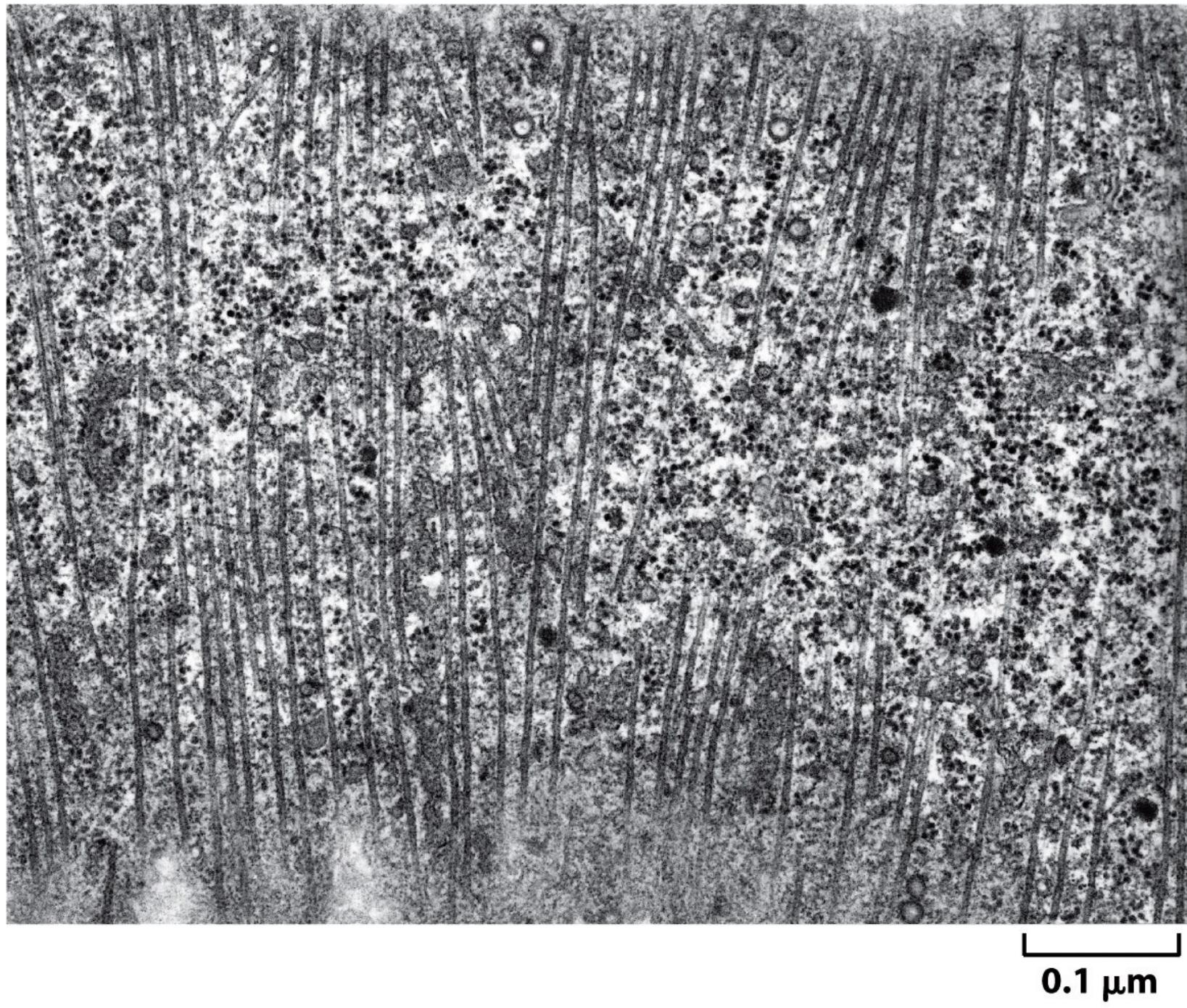


Figure 19-81a *Molecular Biology of the Cell* (© Garland Science 2008)

TIP:

Sintesis es  
extracelular

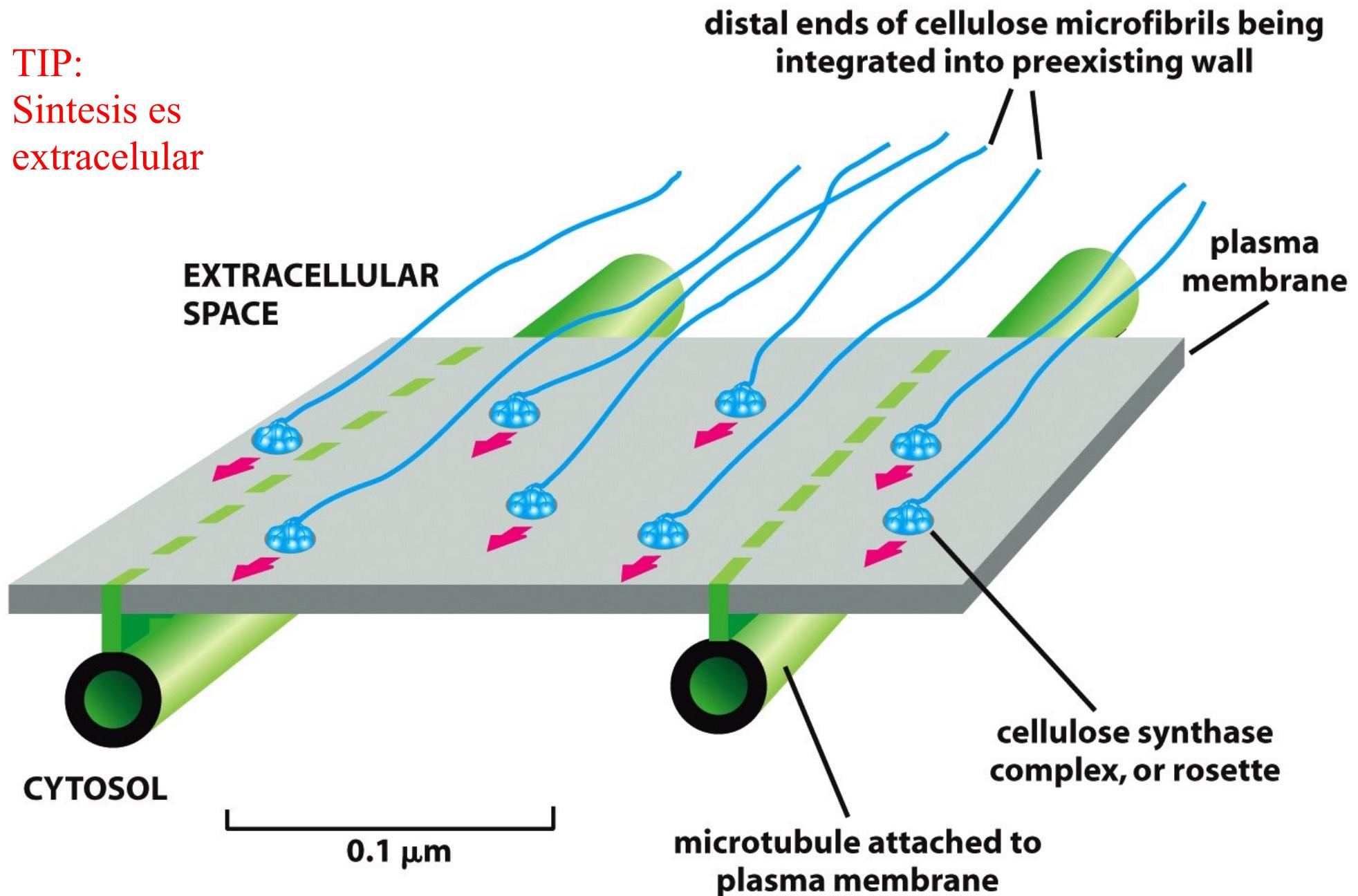


Figure 19-82a *Molecular Biology of the Cell* (© Garland Science 2008)

Lo que vamos a ver....

1.- Matrix extracelular animal y vegetal

2.- Uniones celulares

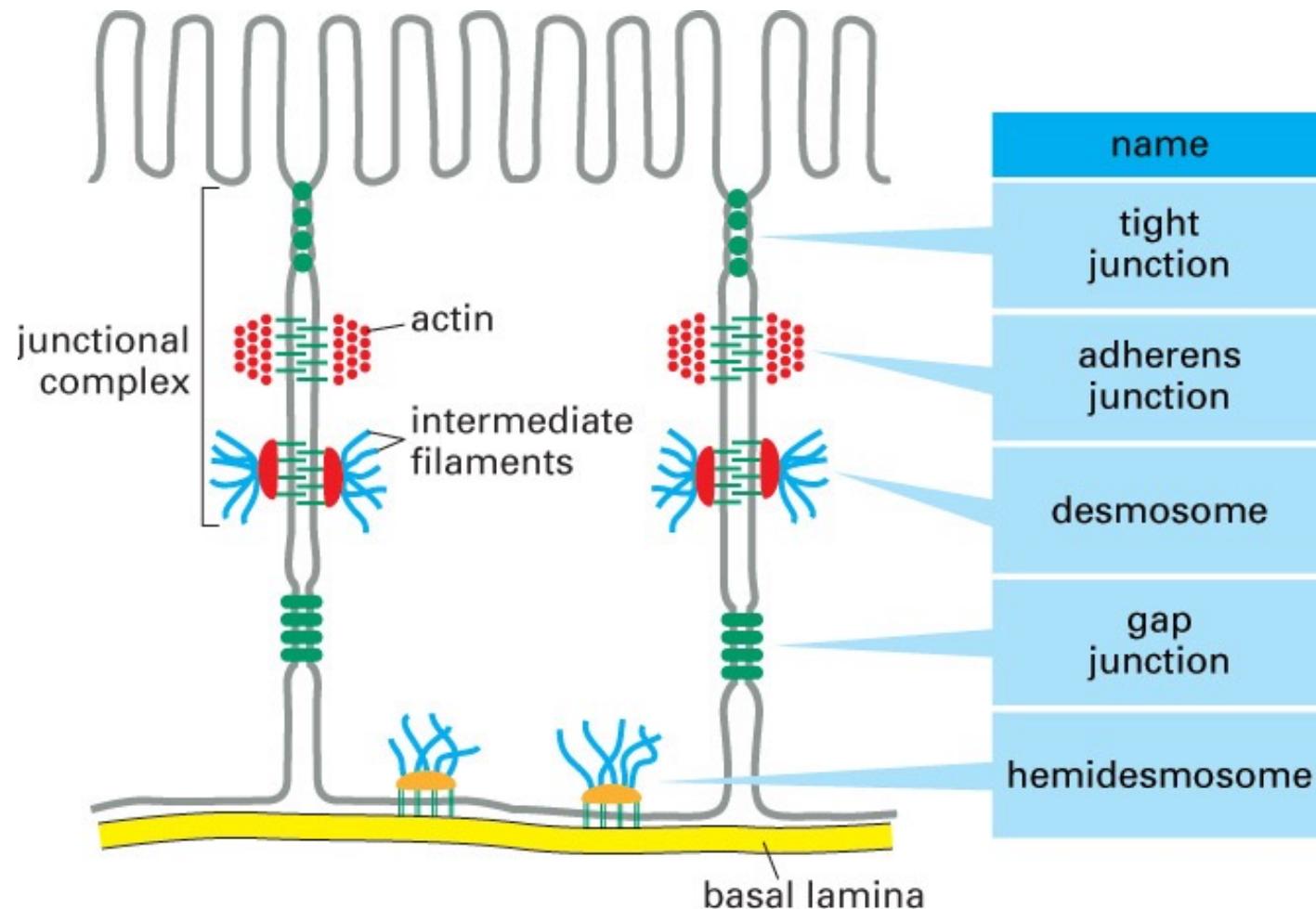
3.- Resumen

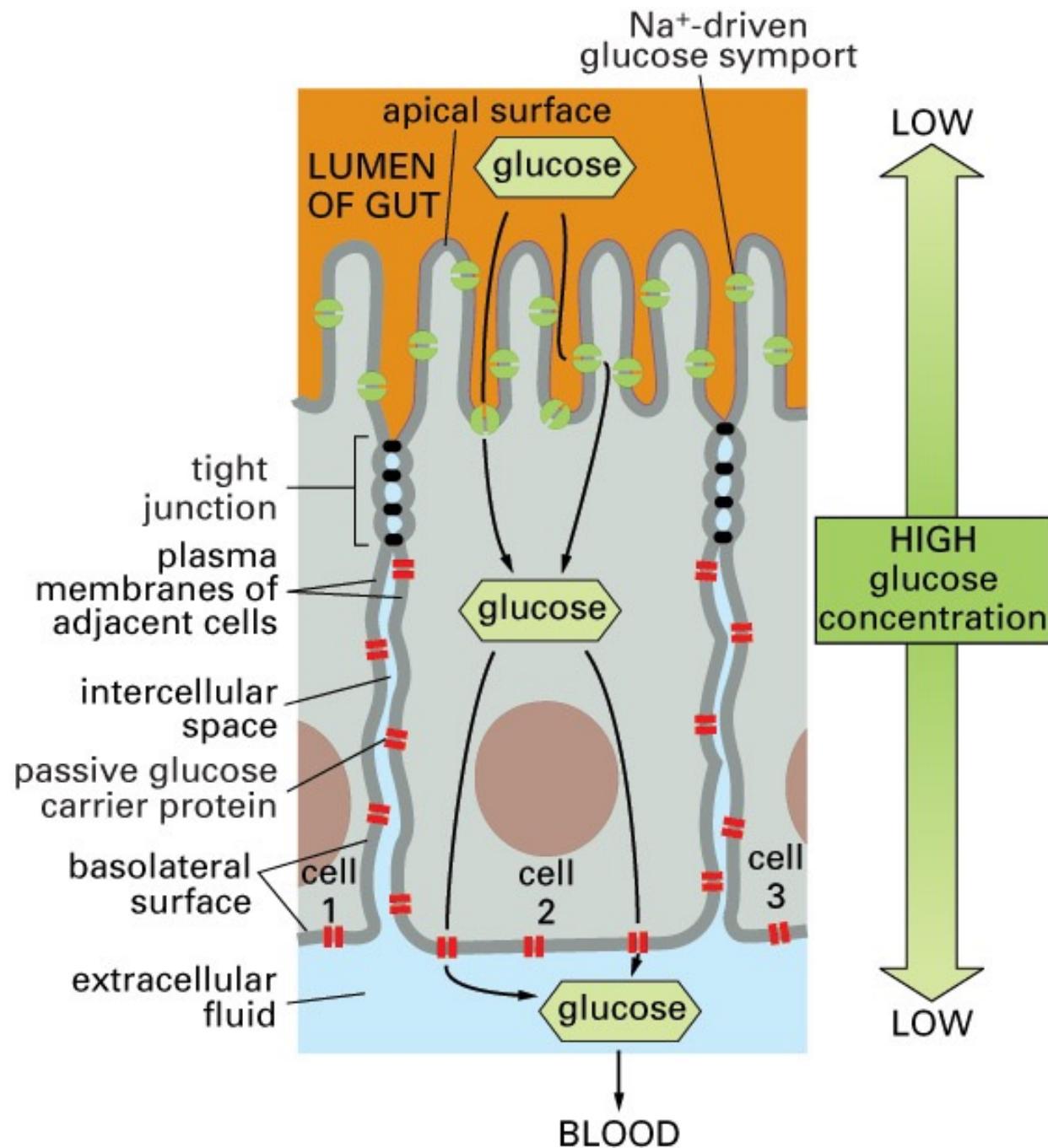
# Tipos de Uniones

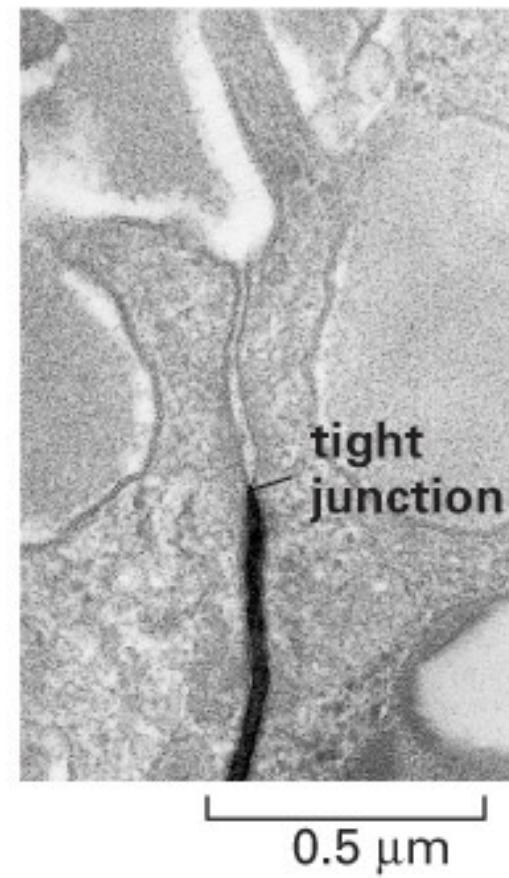
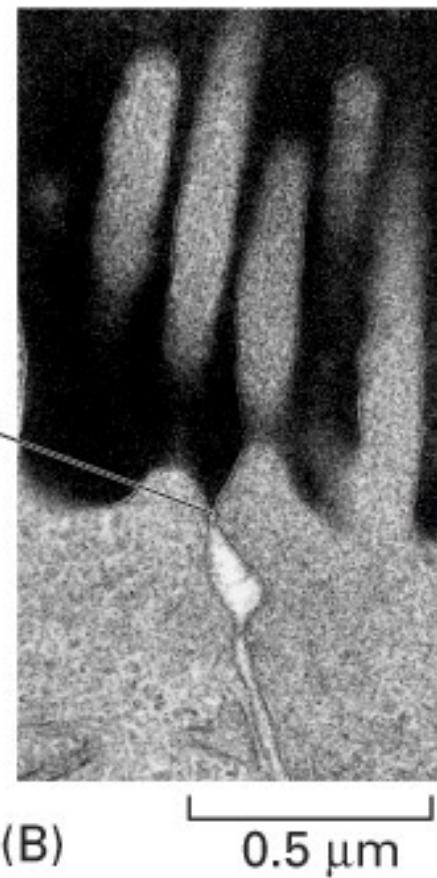
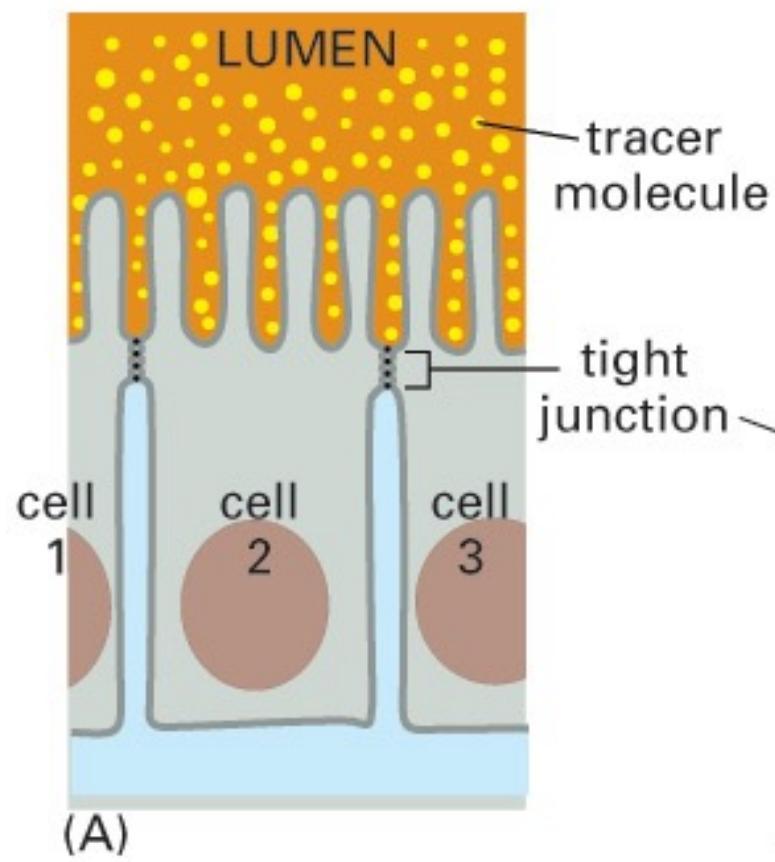
- Uniones Ocludentes:
  - Uniones Estrechas (Tight Junctions)
  - Uniones Septadas (solo en insectos)
- Uniones de Anclaje:
  - Uniones Adherentes (uniones celula-celula)
  - Desmosomas (uniones celula-celula)
  - Hemidesmosomas (uniones celula-MEC)
  - Contactos Focales (uniones celula-MEC)
- Uniones Comunicantes:
  - Uniones en Hendidura (Gap Junctions)
  - Sinápsis químicas
  - Plasmodesmos (solo en plantas)

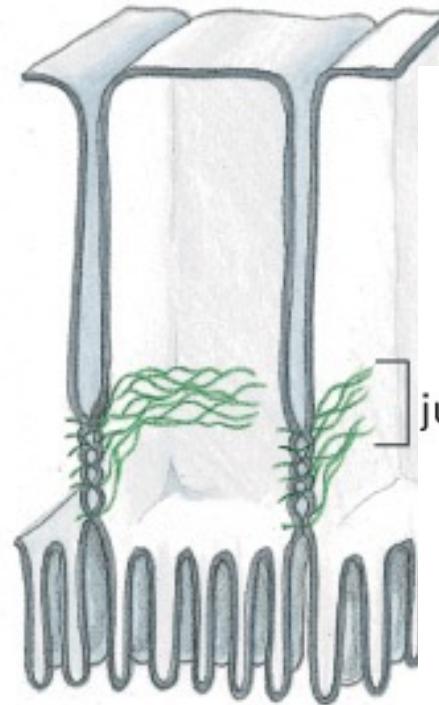
# Tipos de Uniones

- **Uniones Ocludentes:**
  - Uniones Estrechas (Tight Junctions)
  - Uniones Septadas (solo en insectos)
- Uniones de Anclaje:
  - Uniones Adherentes (uniones celula-celula)
  - Desmosomas (uniones celula-celula)
  - Hemidesmosomas (uniones celula-MEC)
  - Contactos Focales (uniones celula-MEC)
- Uniones Comunicantes:
  - Uniones en Hendidura (Gap Junctions)
  - Sinápsis químicas
  - Plasmodesmos (solo en plantas)

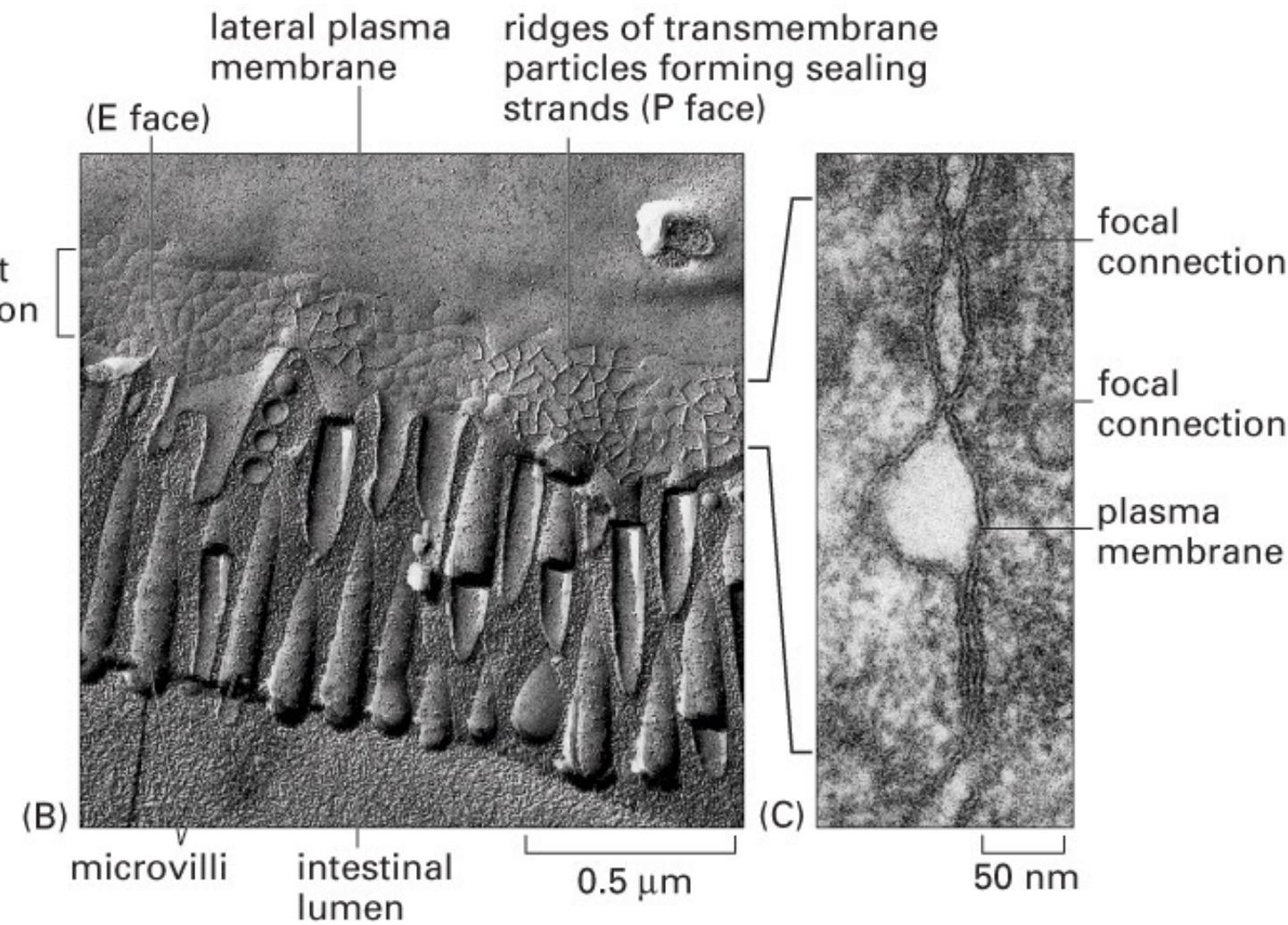


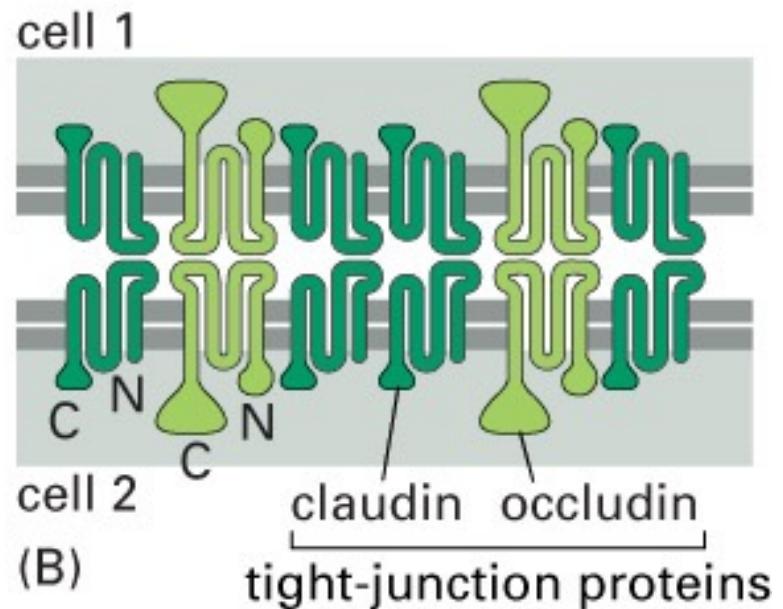
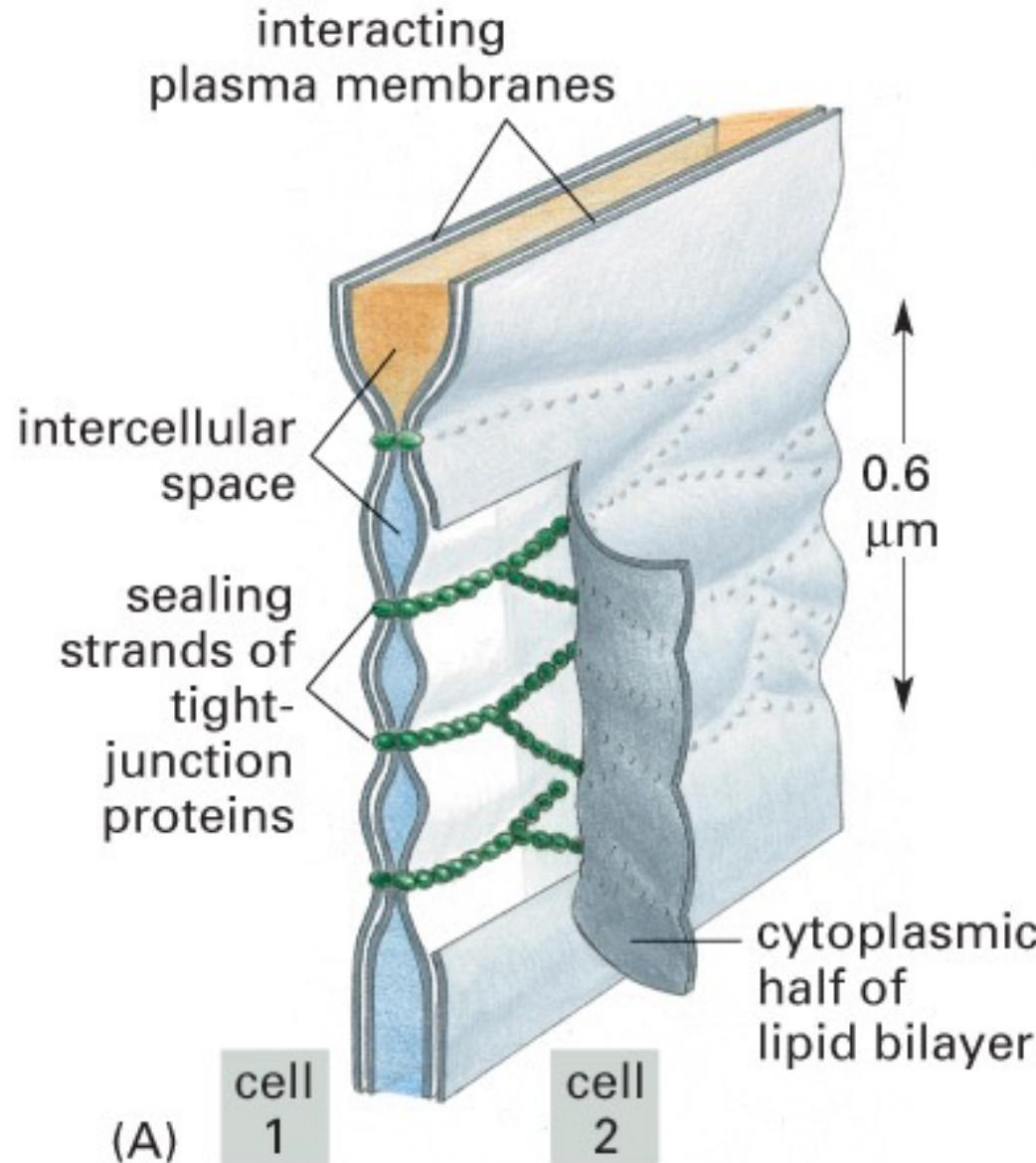






(A)



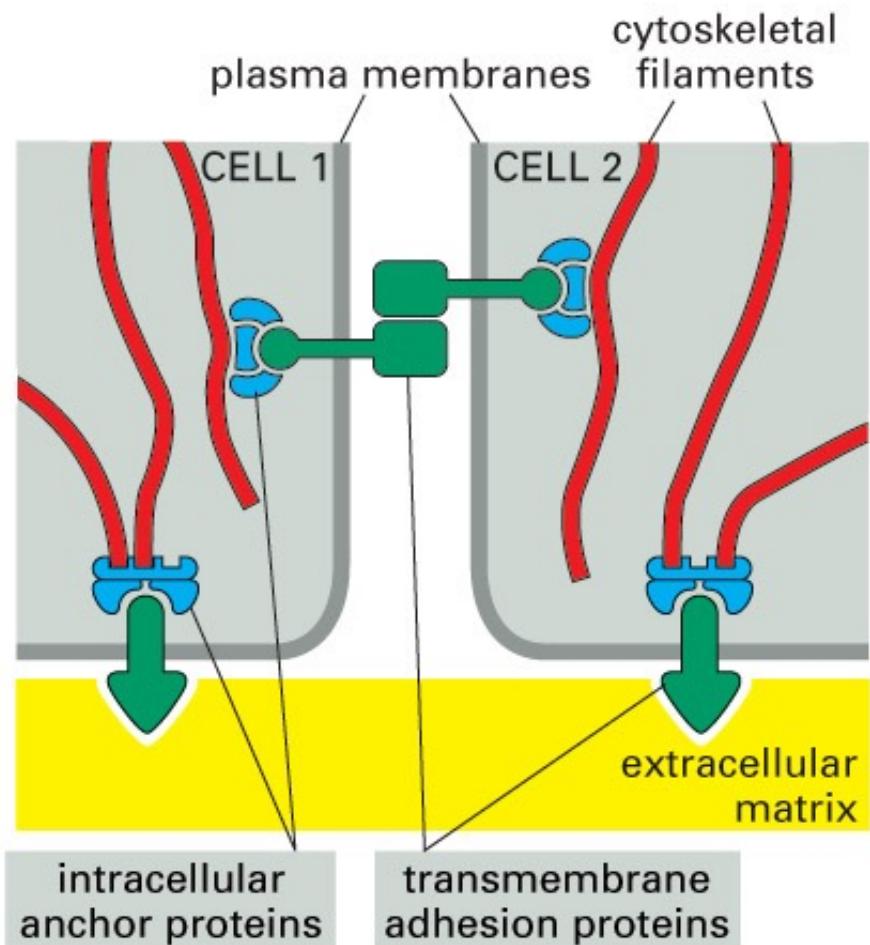
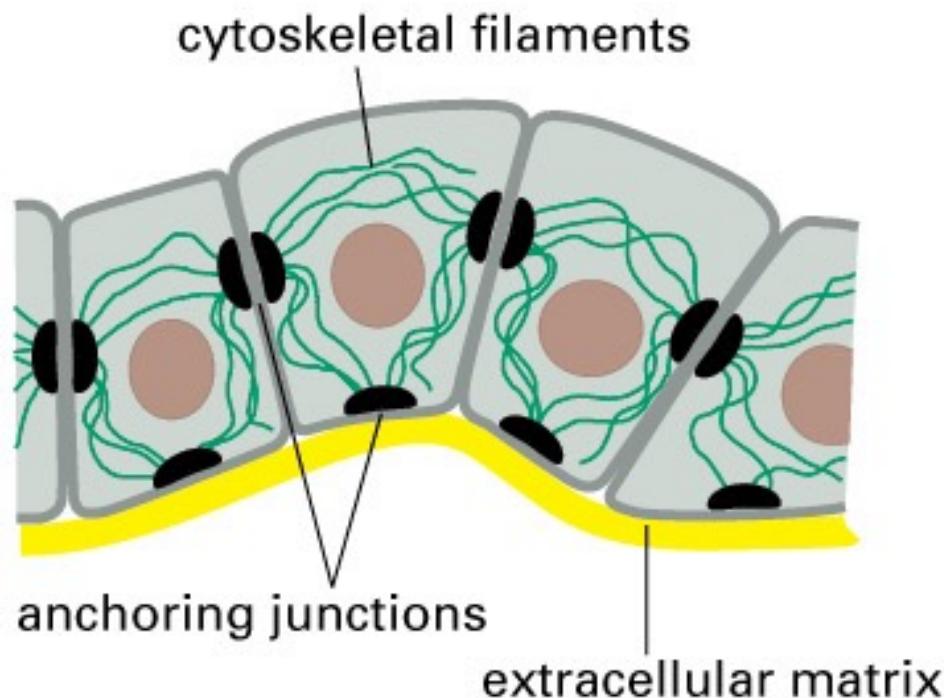


TIP: Proteínas ZO (*zonula occludens*)  
 Invertebrados  
 uniones estrecha:uniones septadas=  
 proteínas ZO:*Disc-large*

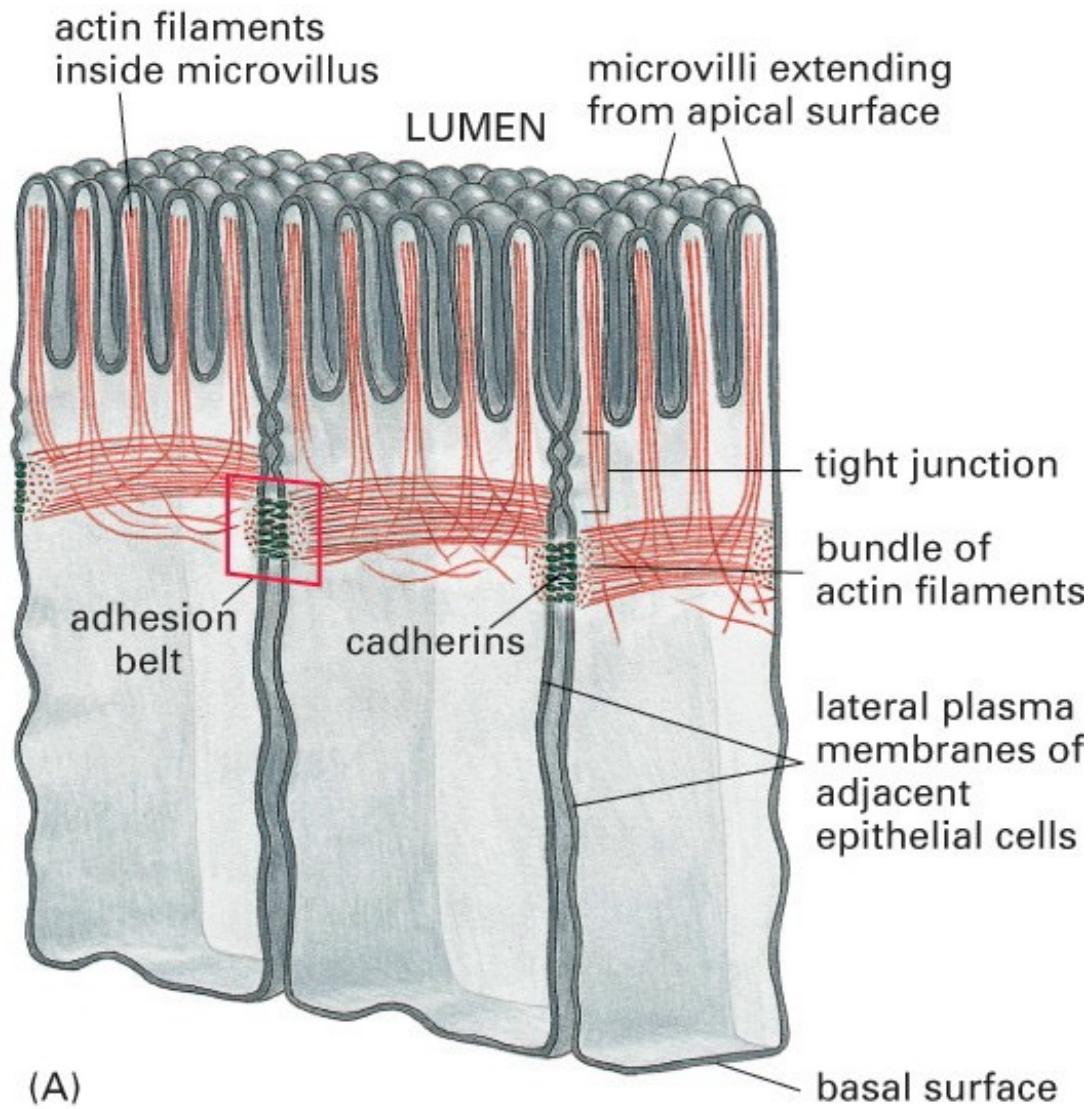
# Tipos de Uniones

- Uniones Ocludentes:
  - Uniones Estrechas (Tight Junctions)
  - Uniones Septadas (solo en insectos)
- Uniones de Anclaje:
  - Uniones Adherentes (uniones celula-celula)
  - Desmosomas (uniones celula-celula)
  - Hemidesmosomas (uniones celula-MEC)
  - Contactos Focales (uniones celula-MEC)
- Uniones Comunicantes:
  - Uniones en Hendidura (Gap Junctions)
  - Sinápsis químicas
  - Plasmodesmos (solo en plantas)

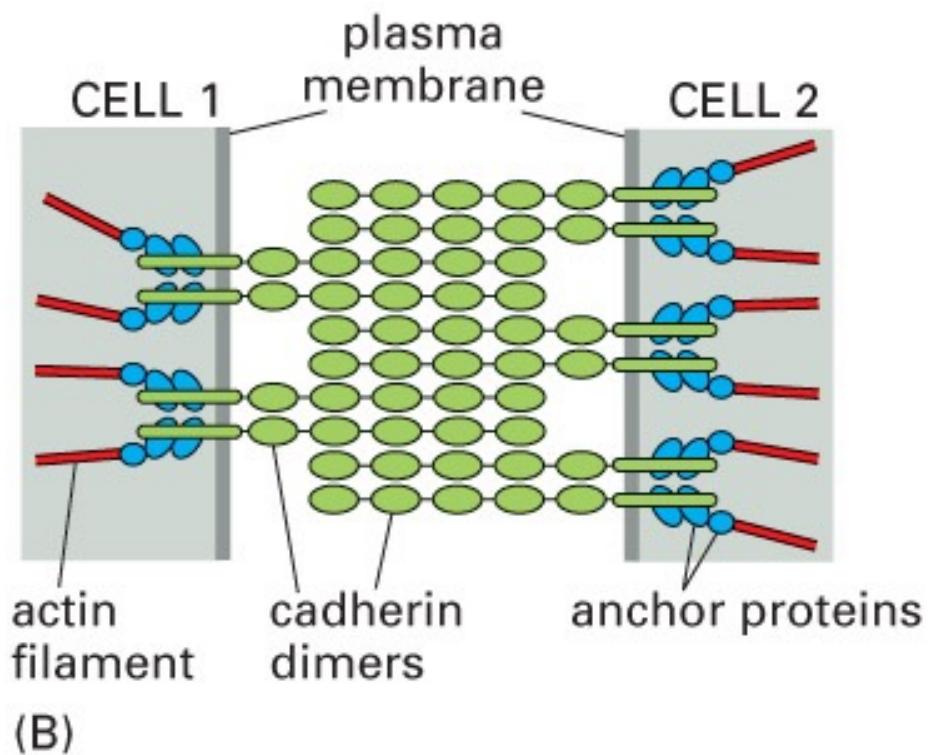
## Uniones de Anclaje



# Uniones Adherentes

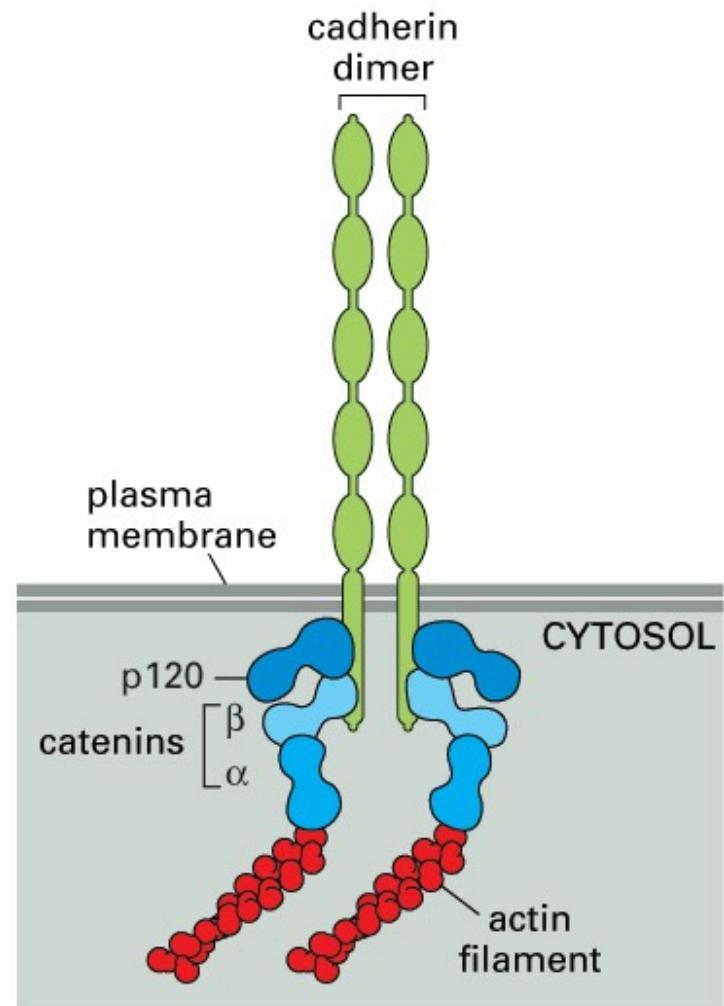
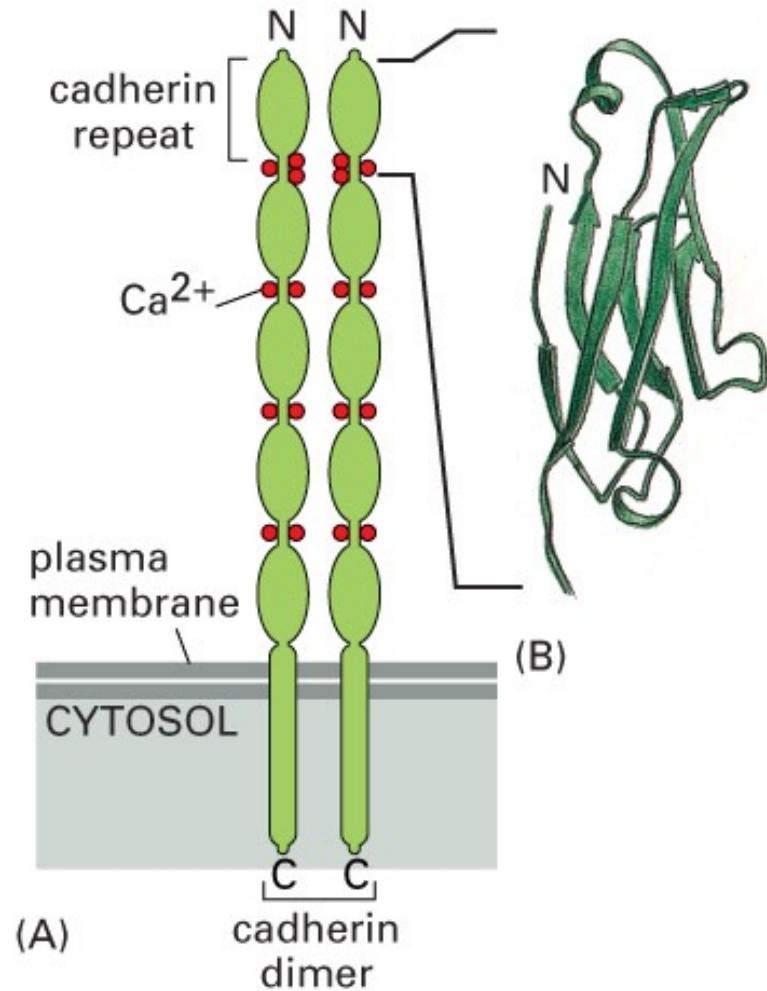


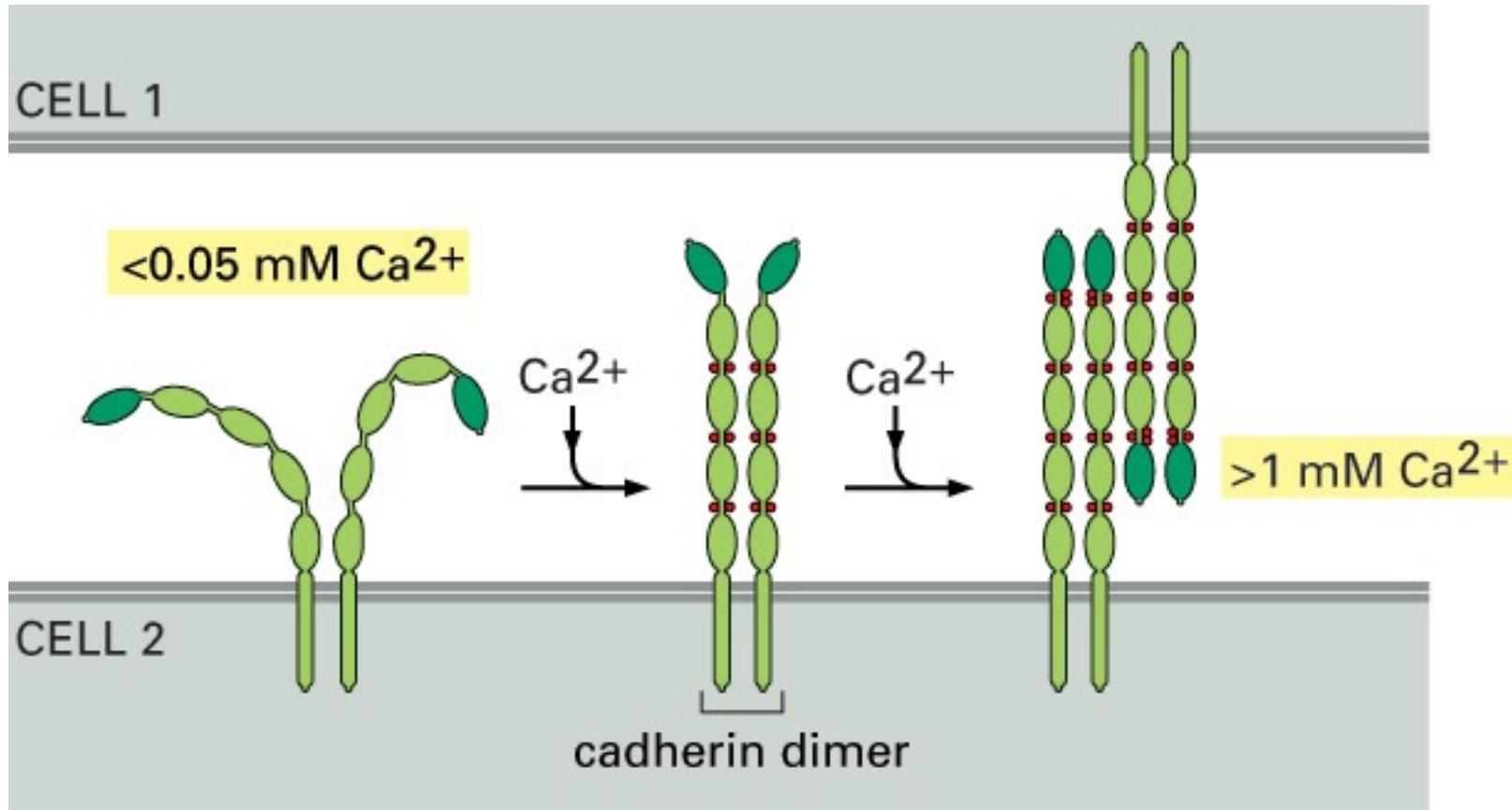
TIP: cinturon de adhesión o *zonula adherens*



(B)

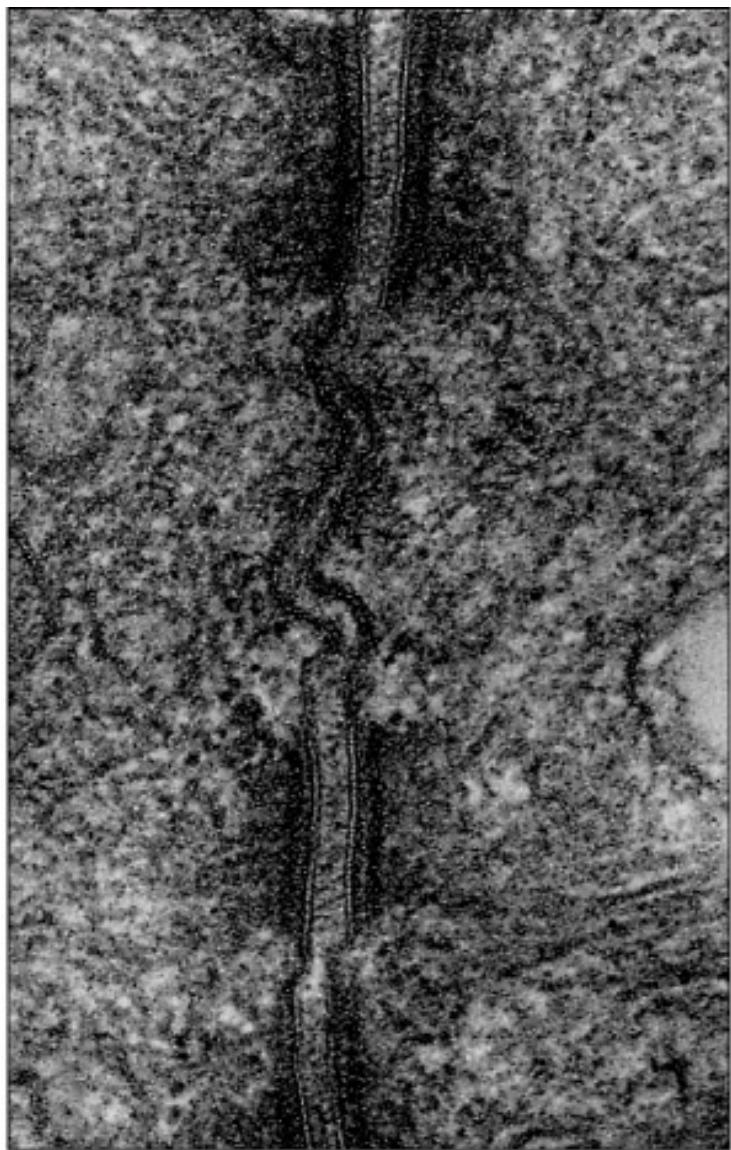
# Cadherinas



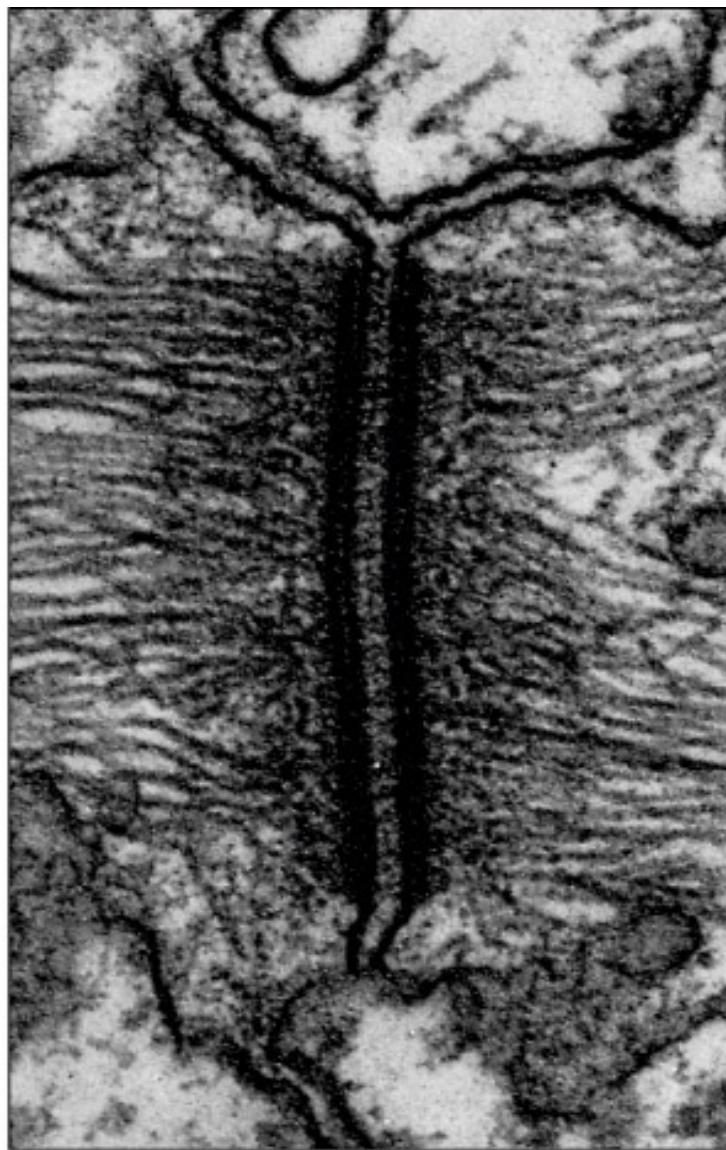


(C)

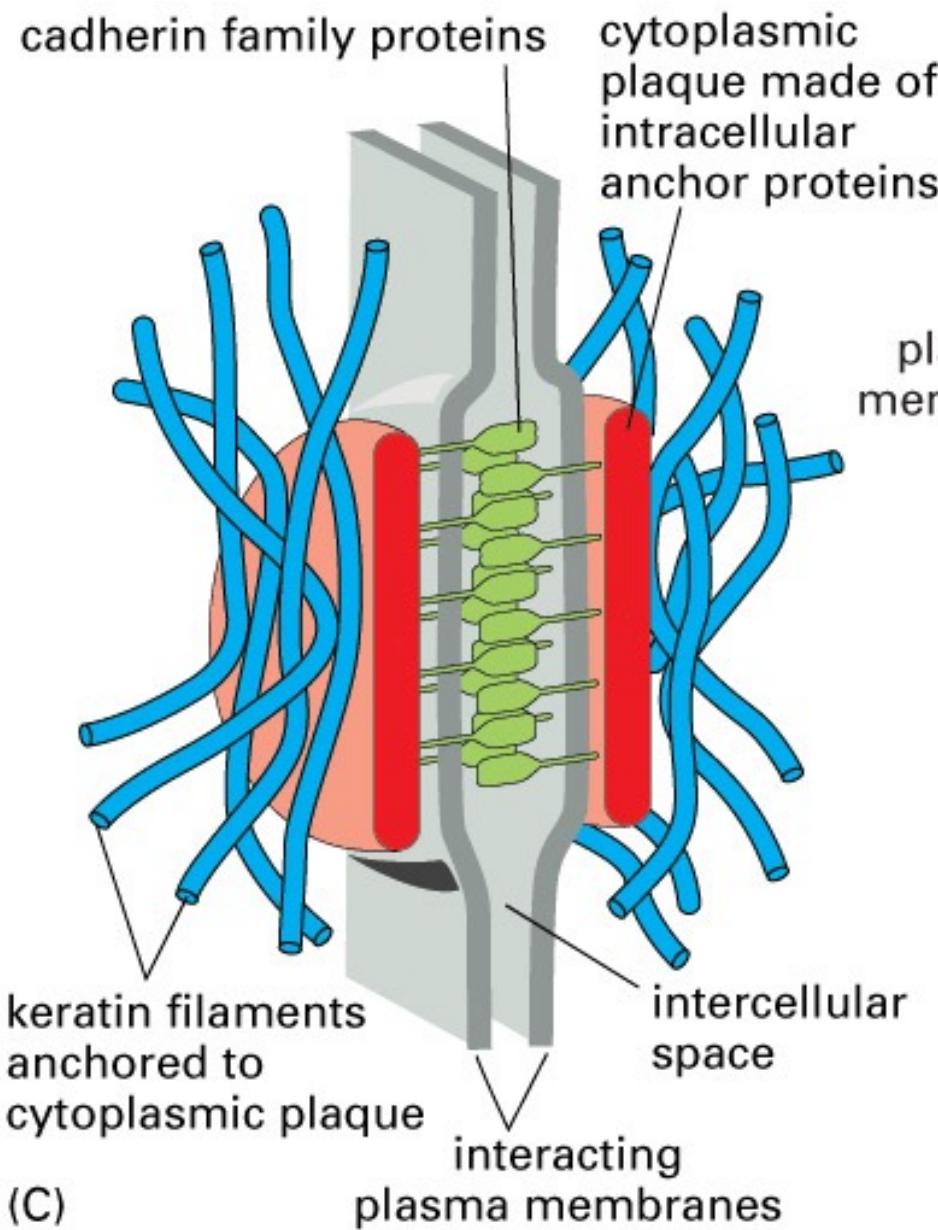
## Desmosomas



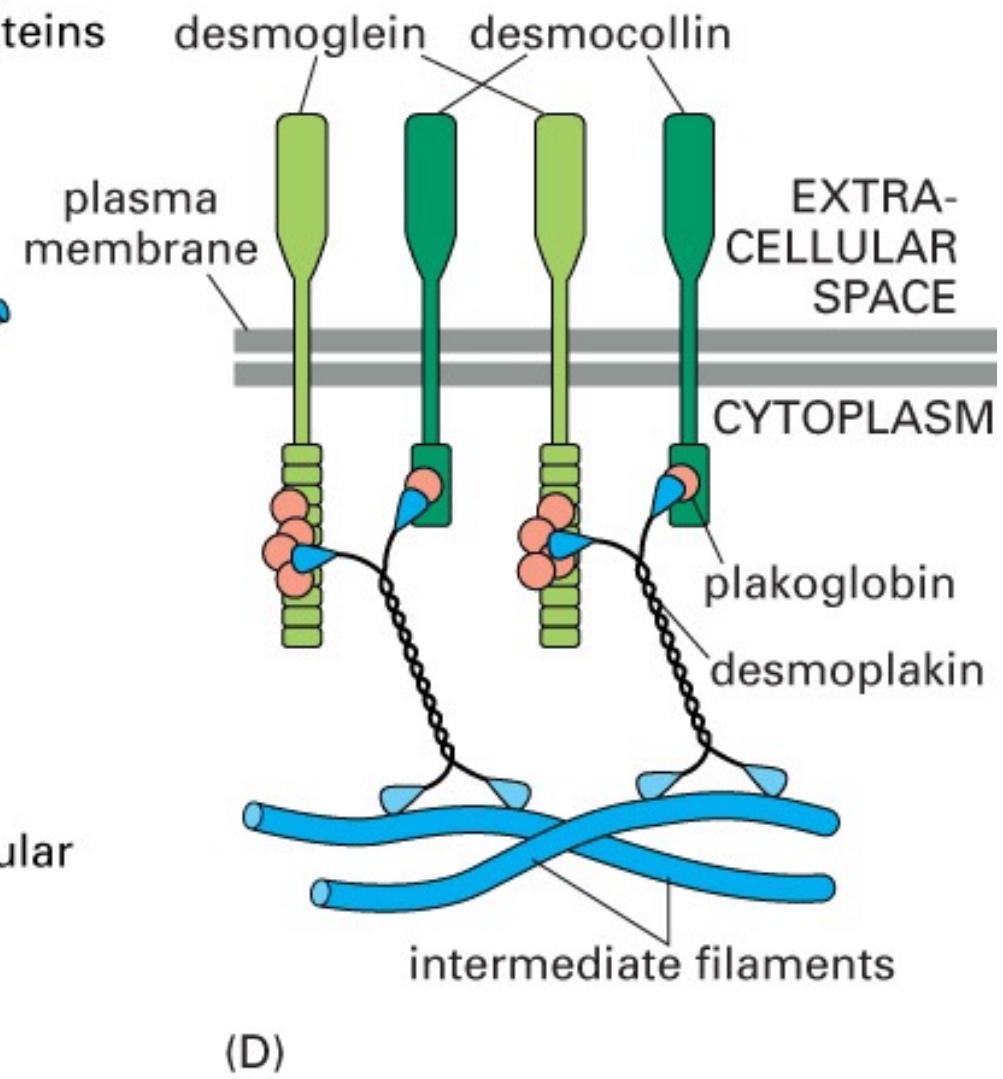
0.1  $\mu\text{m}$



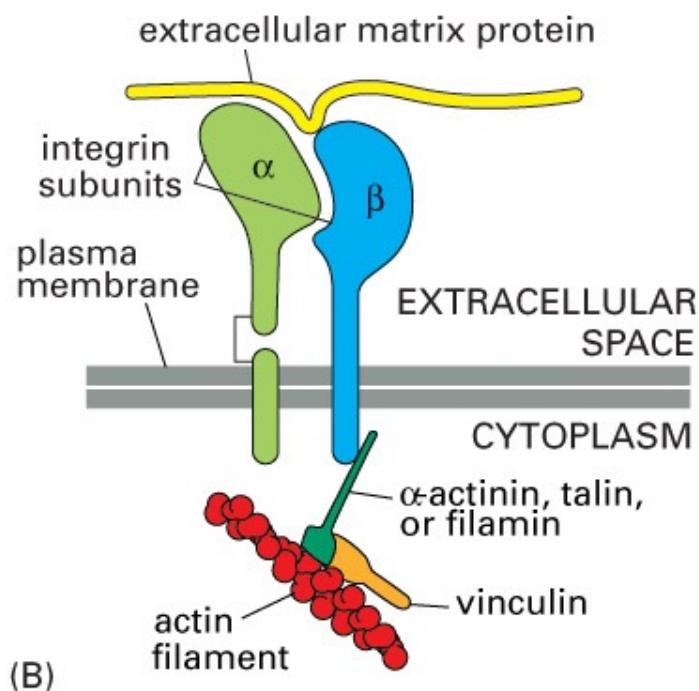
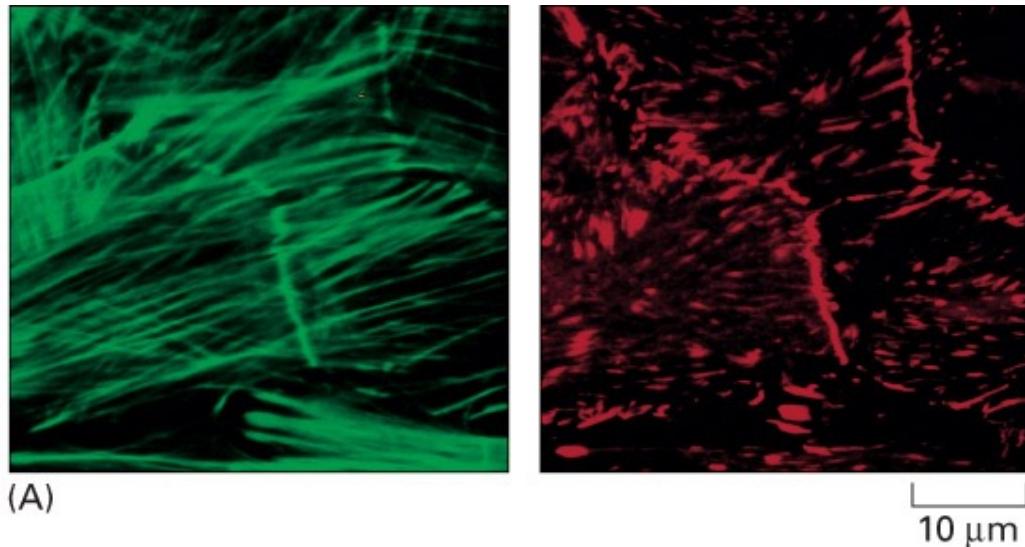
0.1  $\mu\text{m}$



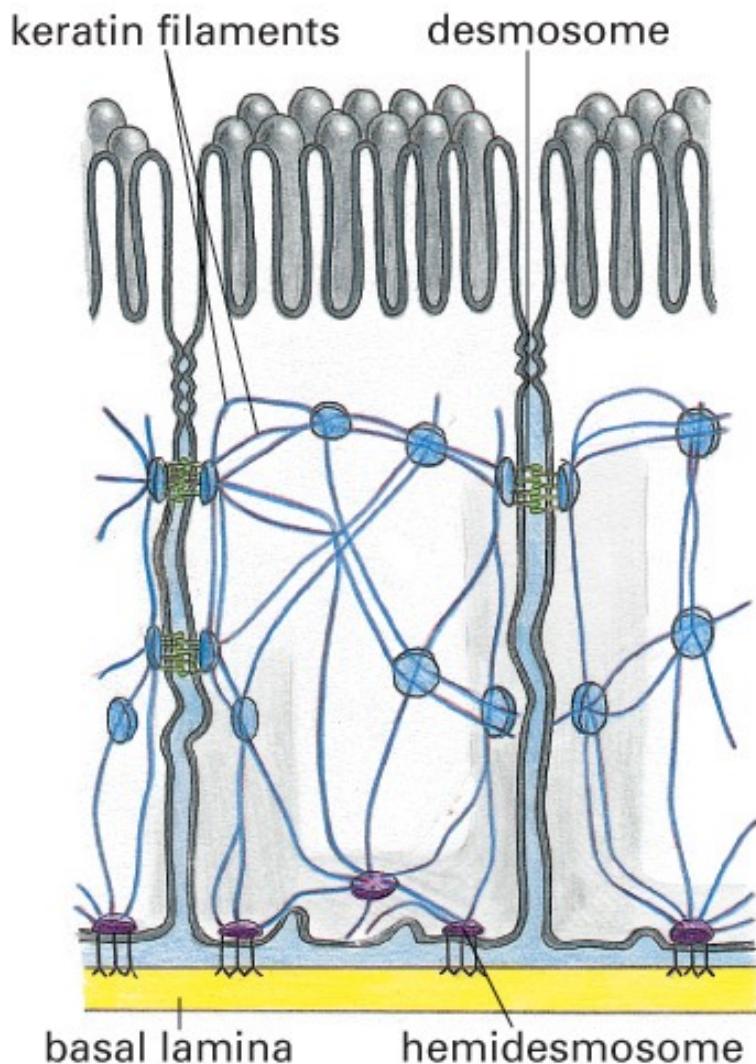
**TIP: desmina (FI) musculo cardiaco**



# Contactos Focales



## Hemidesmosomas

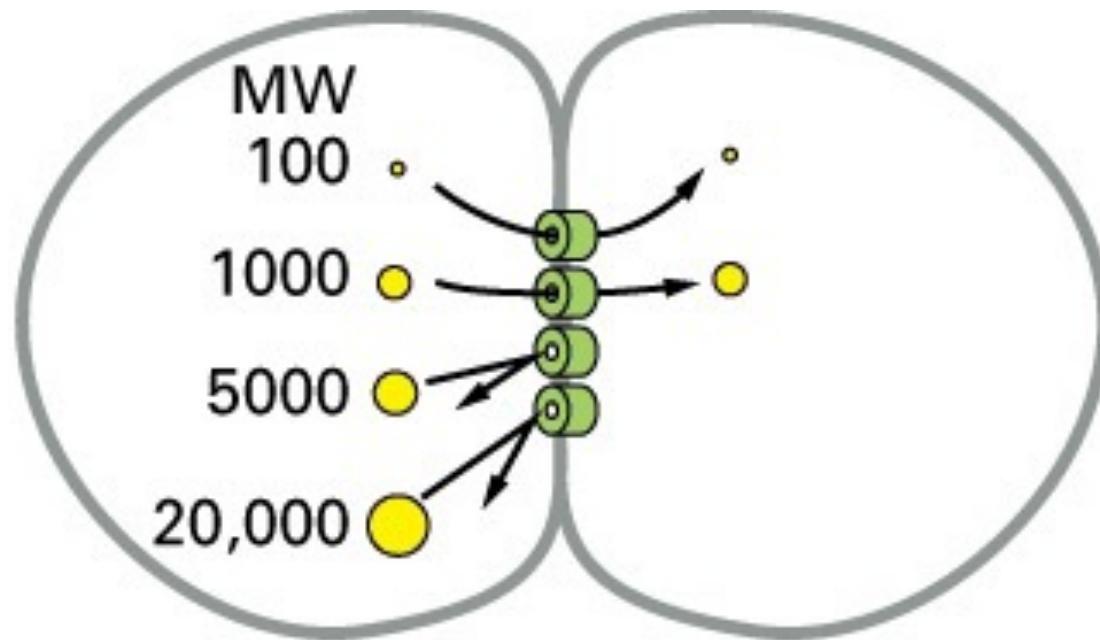


**TIP: Desmosomas (Desmocolina, desmogleinas; Placoglobinas, Desmoplaquinas: Filamentos Intermedios).**  
**Hemidesmosomas (Laminina: Integrinas: Plectinas:Filamentos intermedios)**

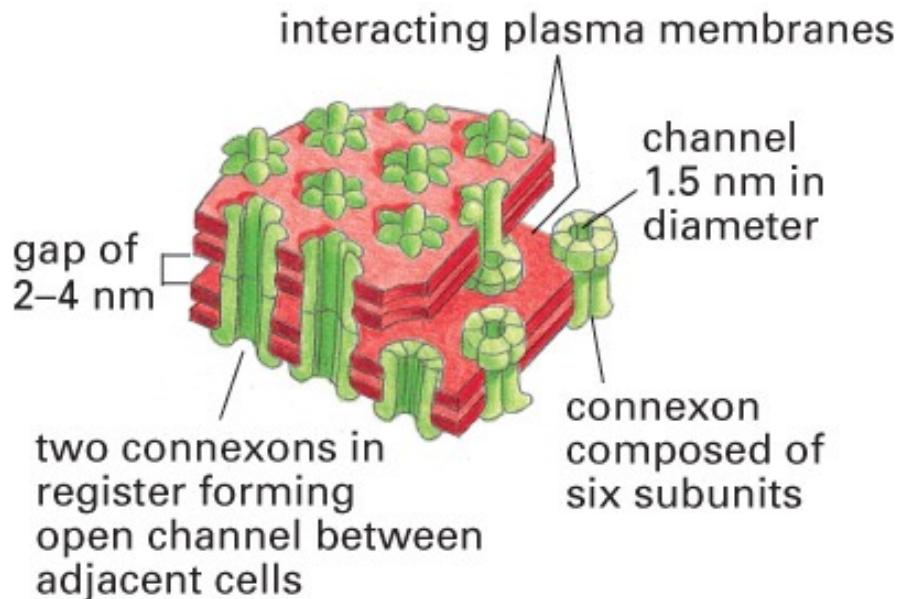
# Tipos de Uniones

- Uniones Ocludentes:
  - Uniones Estrechas (Tight Junctions)
  - Uniones Septadas (solo en insectos)
- Uniones de Anclaje:
  - Uniones Adherentes (uniones celula-celula)
  - Desmosomas (uniones celula-celula)
  - Hemidesmosomas (uniones celula-MEC)
  - Contactos Focales (uniones celula-MEC)
- Uniones Comunicantes:
  - Uniones en Hendidura (Gap Junctions)
  - Plasmodesmos (solo en plantas)

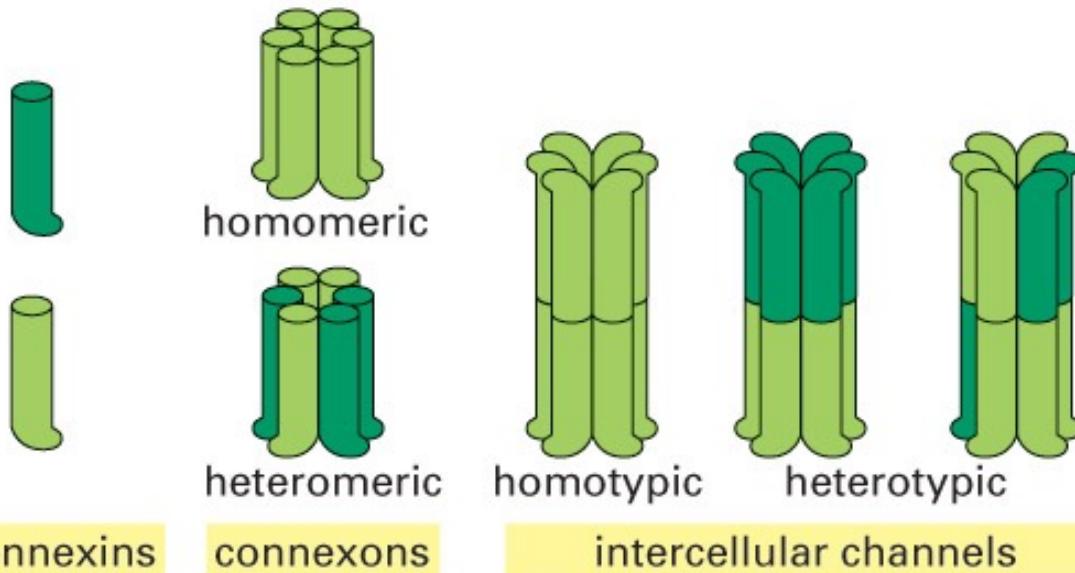
## Uniones en Hendidura



TIP: iones, azucares, aminoacidos, nucleotidos, cAMP e IP3)



(A)



(B)



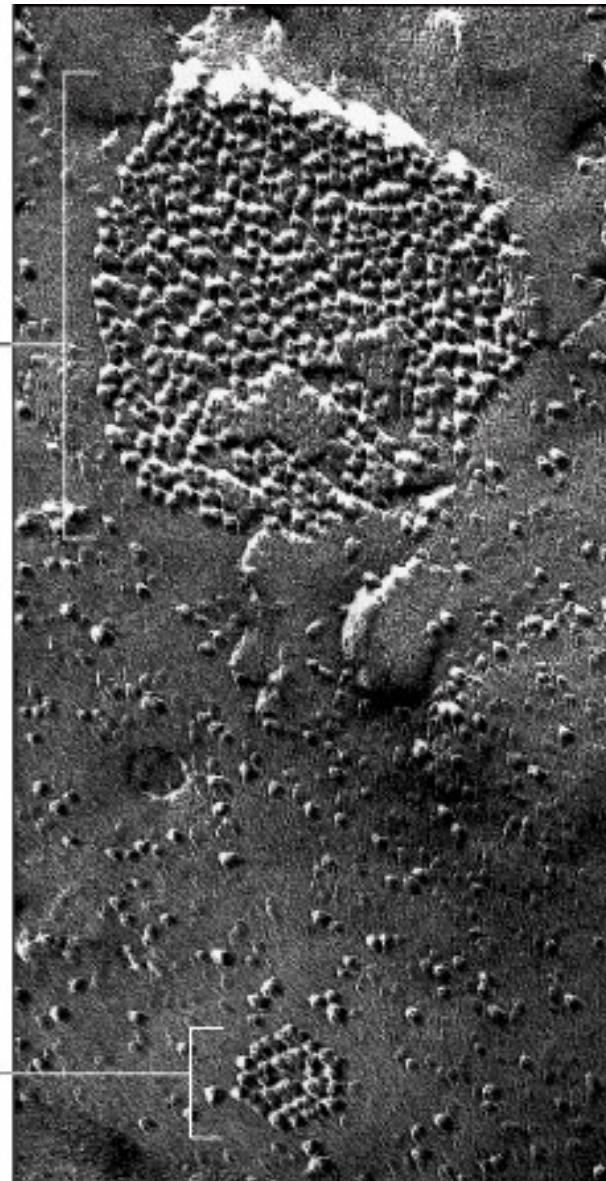
(A)

100 nm

large  
gap  
junction

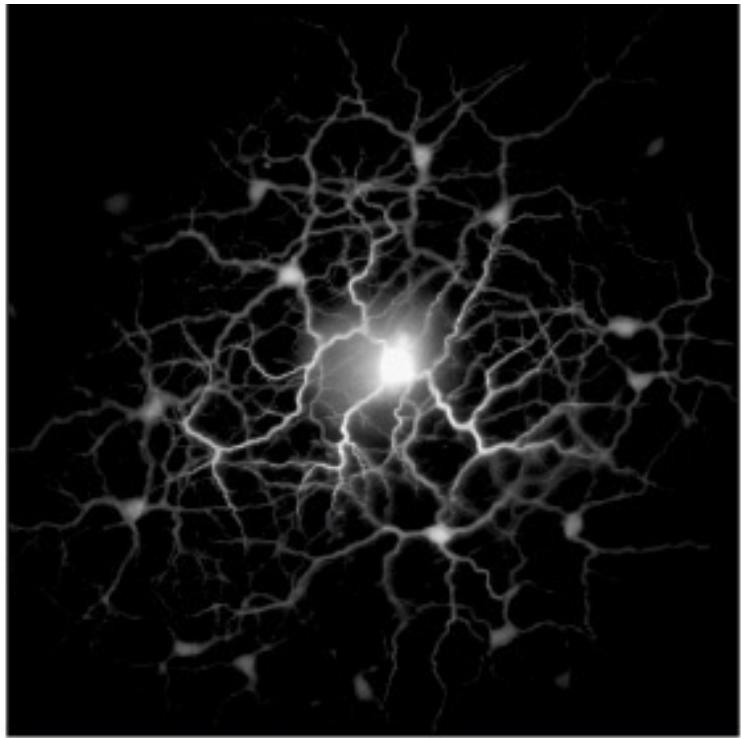
membranes

small  
gap  
junction

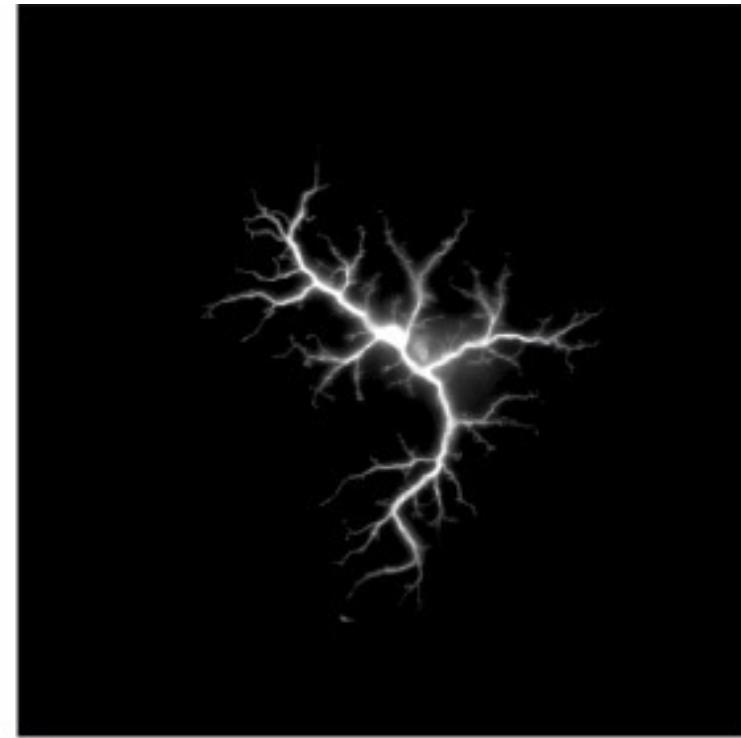


(B)

100 nm

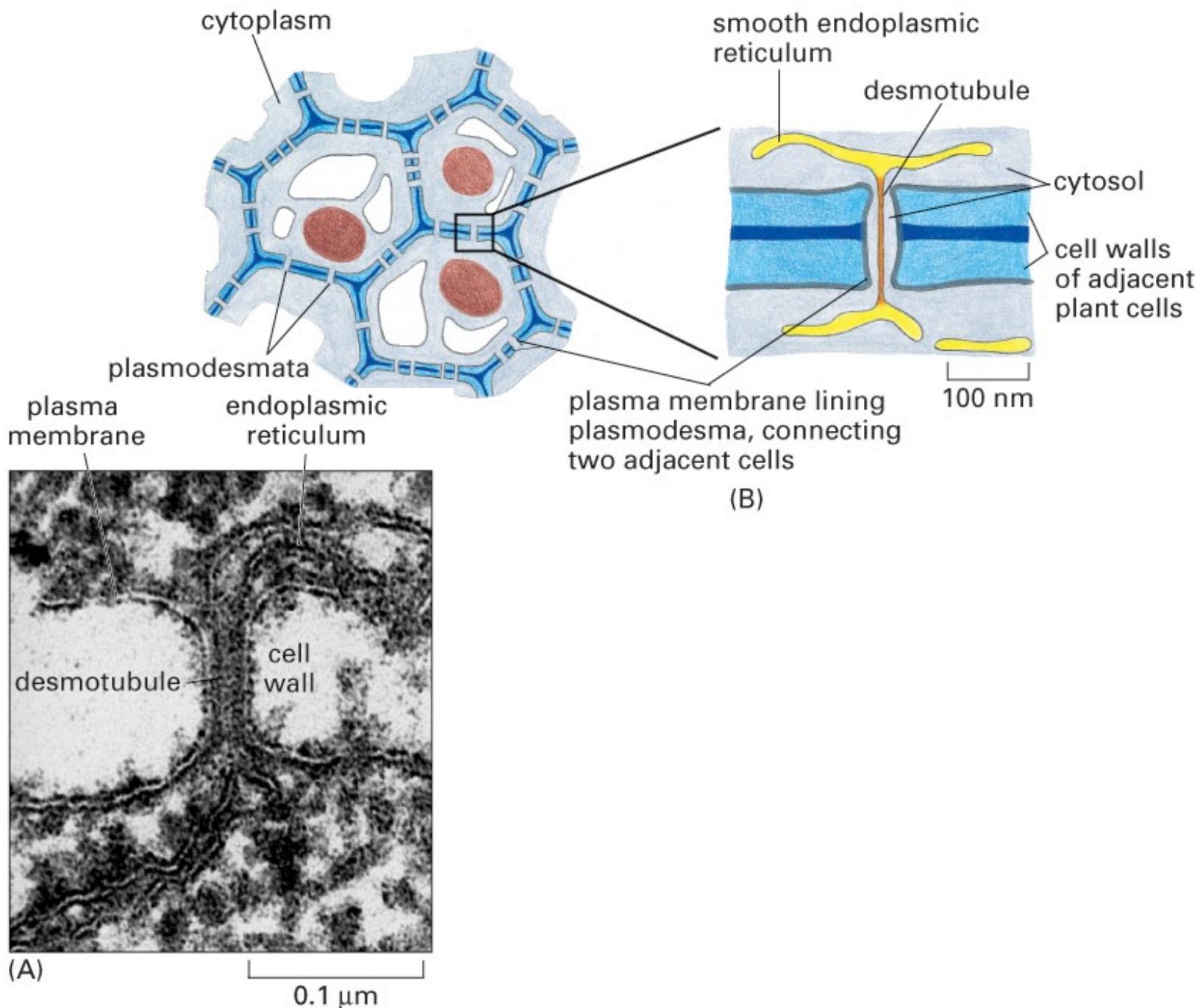


(A)



(B)

# Plasmodesmos

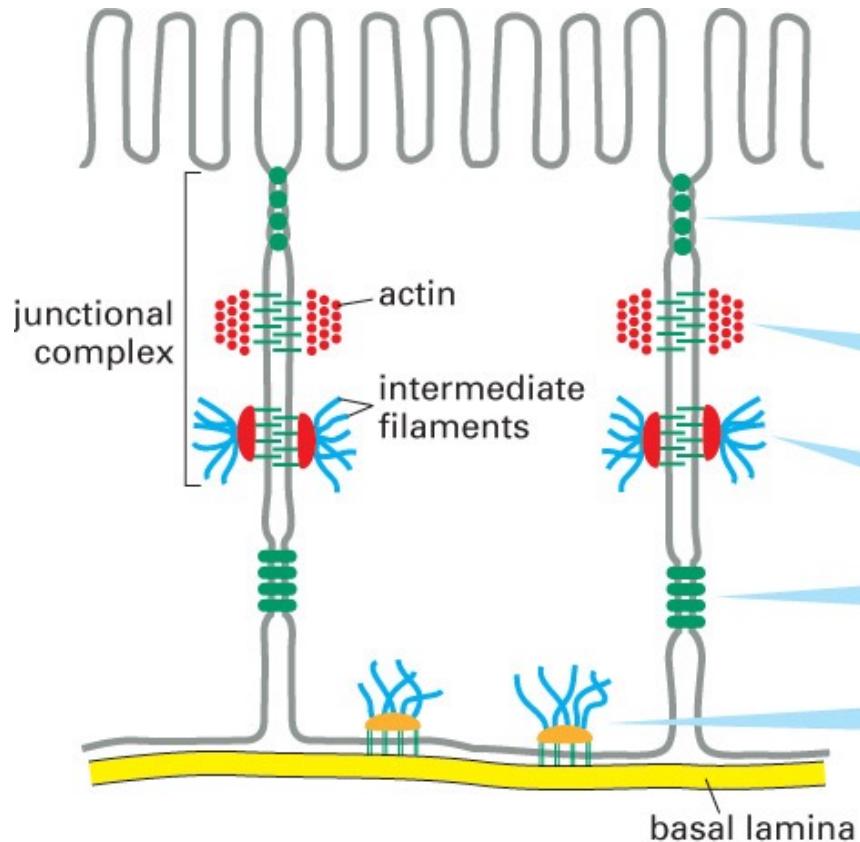


Lo que vamos a ver....

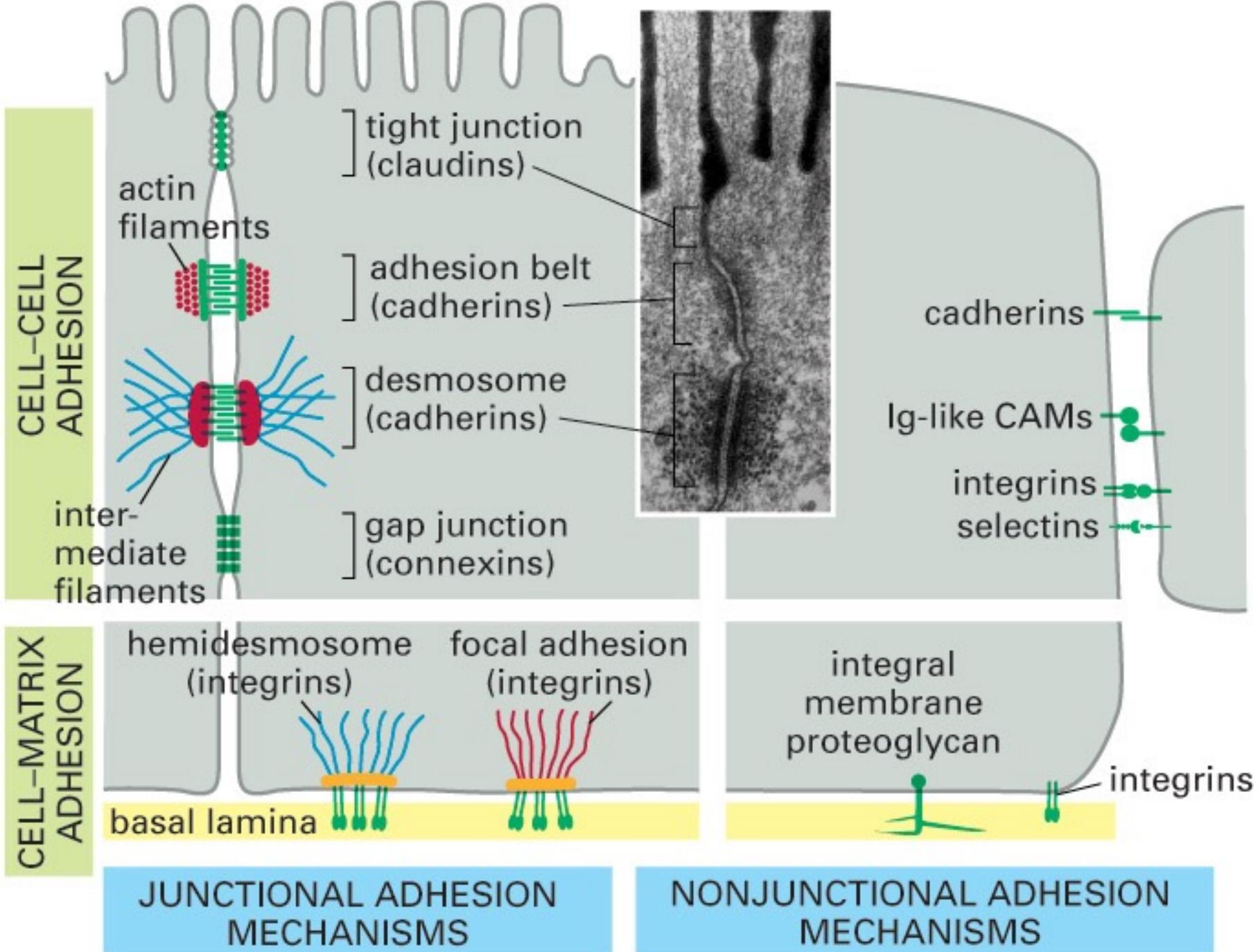
1.- Matrix extracelular animal y vegetal

2.- Uniones celulares

3.- Resumen



name	function
tight junction	seals neighboring cells together in an epithelial sheet to prevent leakage of molecules between them
adherens junction	joins an actin bundle in one cell to a similar bundle in a neighboring cell
desmosome	joins the intermediate filaments in one cell to those in a neighbor
gap junction	allows the passage of small water-soluble ions and molecules
hemidesmosome	anchors intermediate filaments in a cell to the basal lamina



## **Bibliografía Clase**

- Molecular Biology of the Cell (Bruce Alberts), 5<sup>th</sup> Edition (2008)**
- Essential Cell Biology (Bruce Alberts), 3<sup>nd</sup> Edition (2010)**