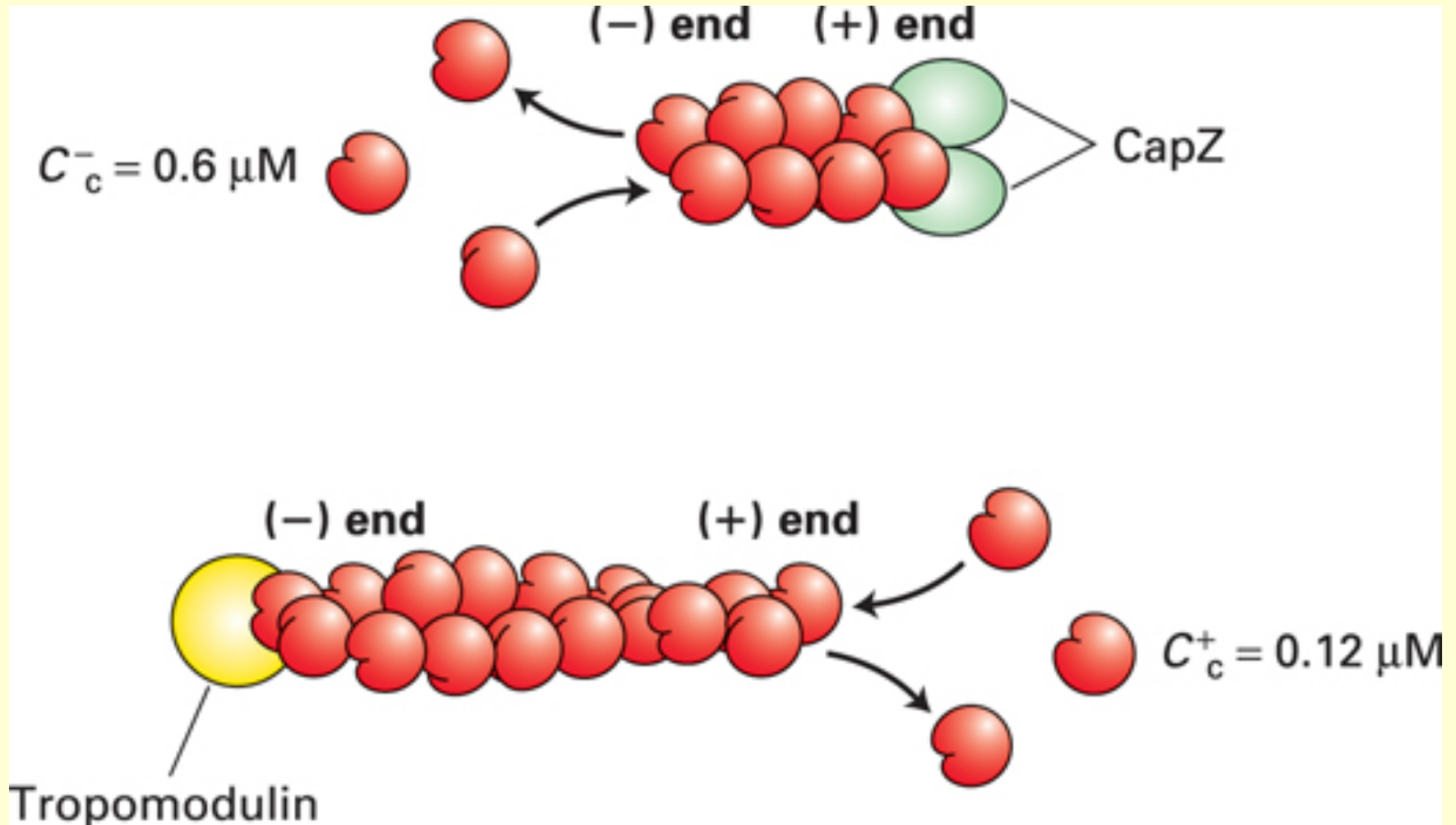
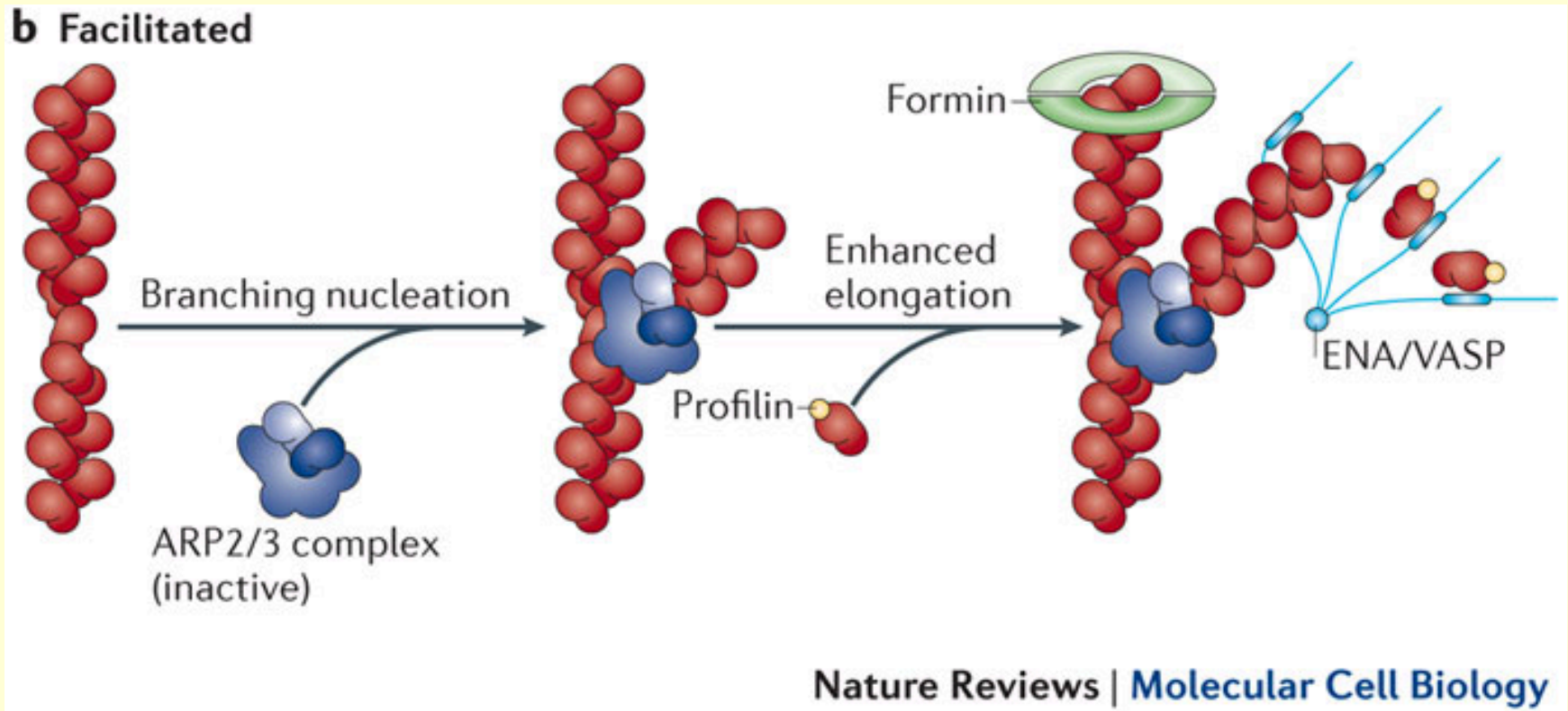


Actin capping proteins **CapZ** and **tropomodulin** bind selectively to either end of a filament to stabilise the filament



Actin filament assembly in the cell - how are new filaments started ?

Actin nucleating proteins *formin* and *Arp2/3*



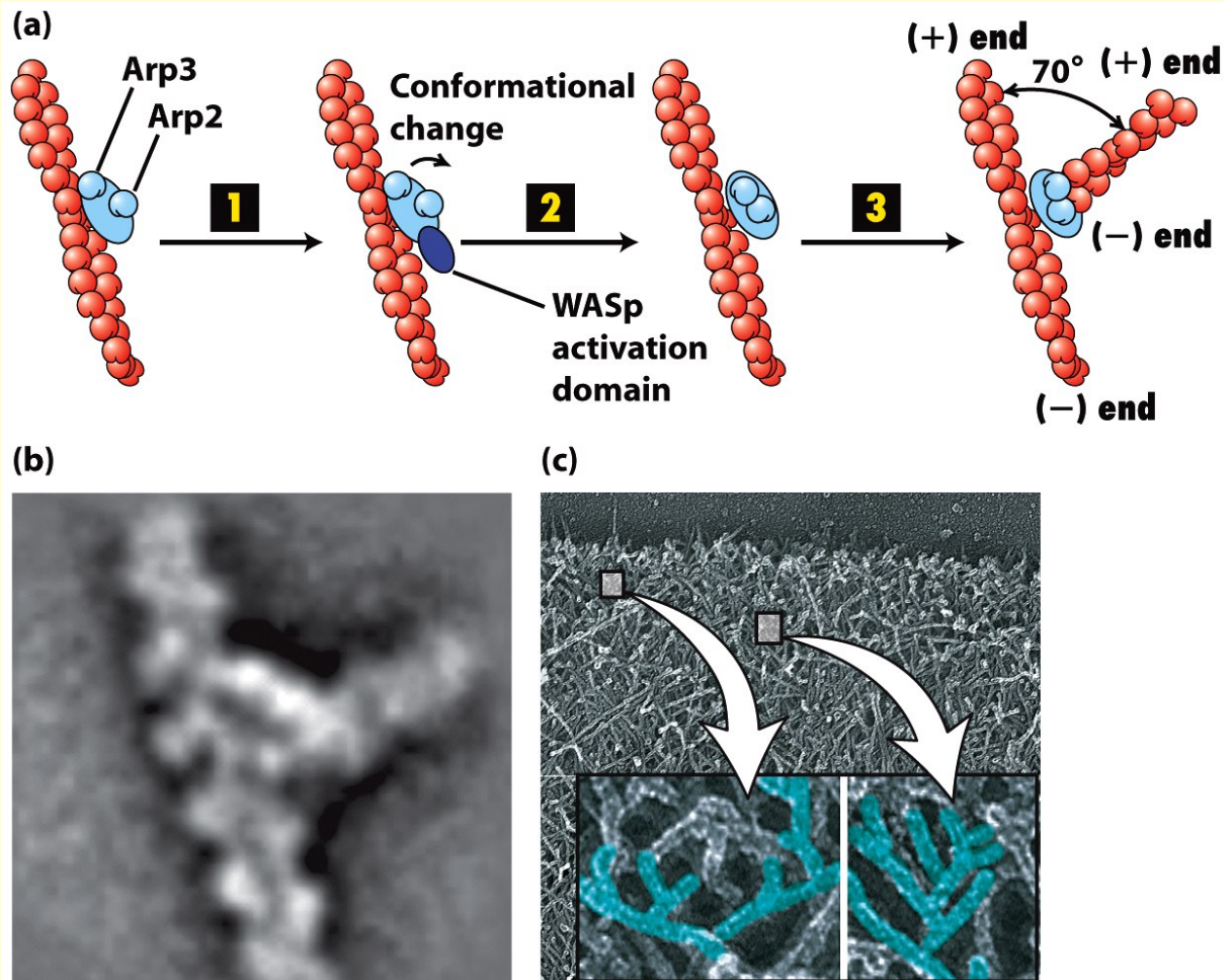
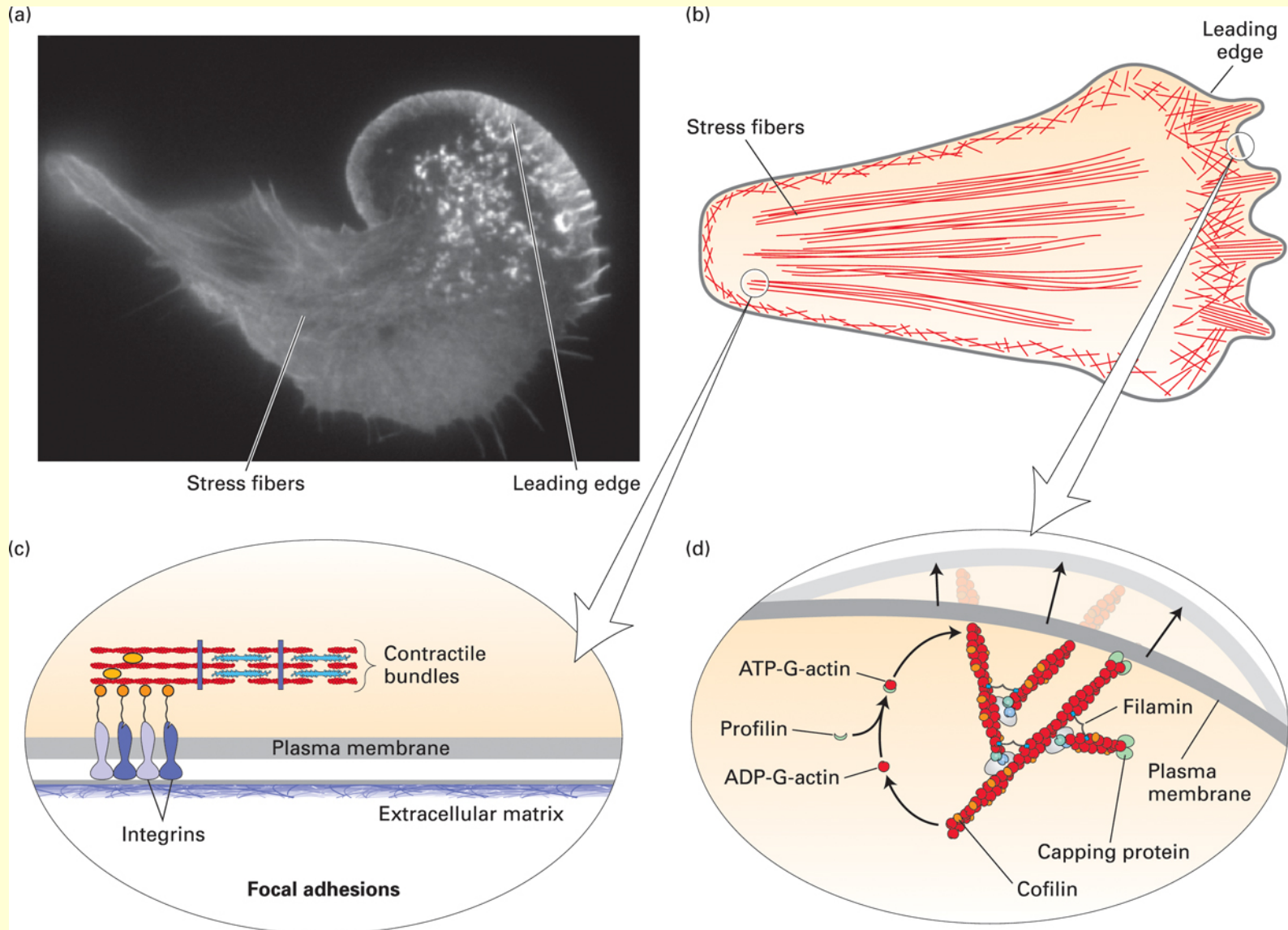


Figure 17-15  
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Branching of actin filaments is regulated by Arp2/3 & WASp

**Figure 17.41 Actin-based structures involved in cell locomotion.**





## **Intramural** Research Program

*Our Research Changes Lives*

# Review of actin-binding proteins and their function

Profilin

Cofilin

CapZ

tropomodulin

Firmin

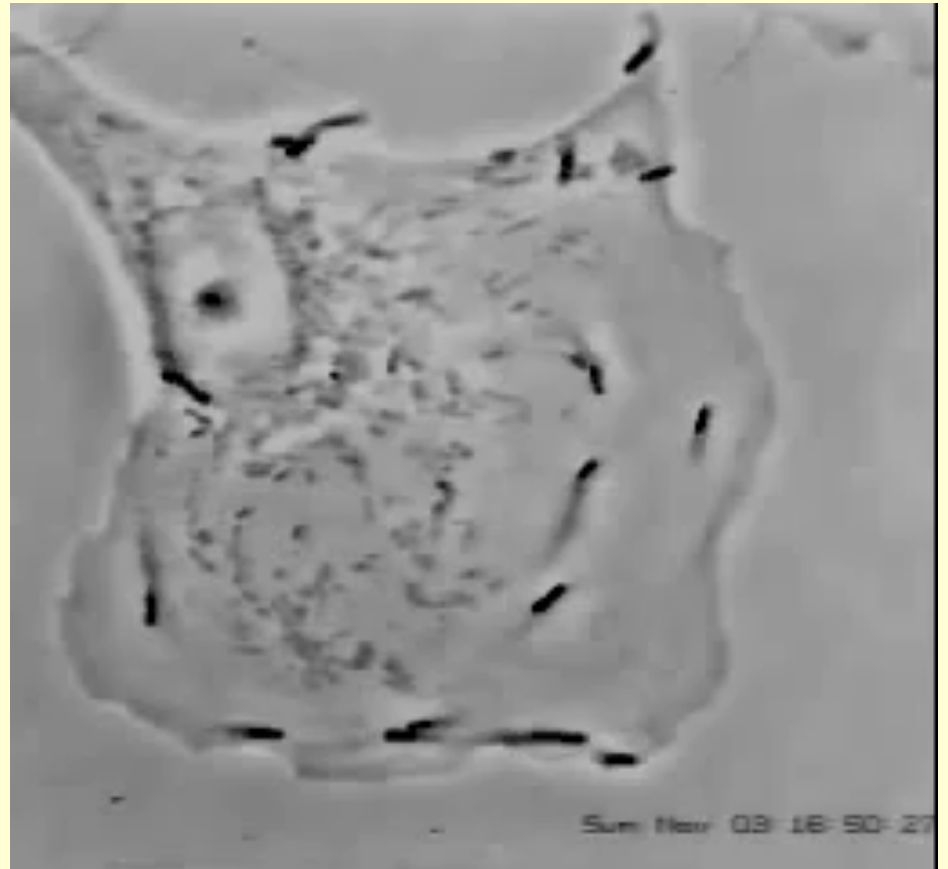
Arp2/3

Learn how each protein interacts with actin and the effect of filament growth



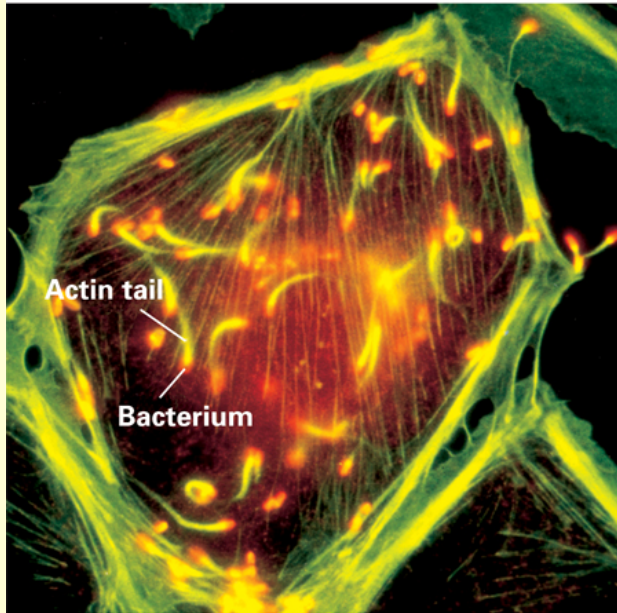
#### 4. Intracellular movement can be powered by actin polymerisation

*Listeria monocytogenes*, a human pathogenic bacteria uses actin polymerisation to push itself around inside a cell and into a neighbouring cell

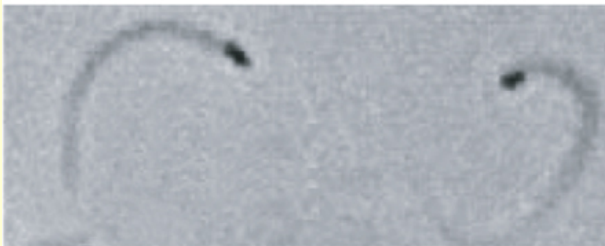


**Experimental Figure 17.17** *Listeria* utilizes the power of actin polymerization for intracellular movement.

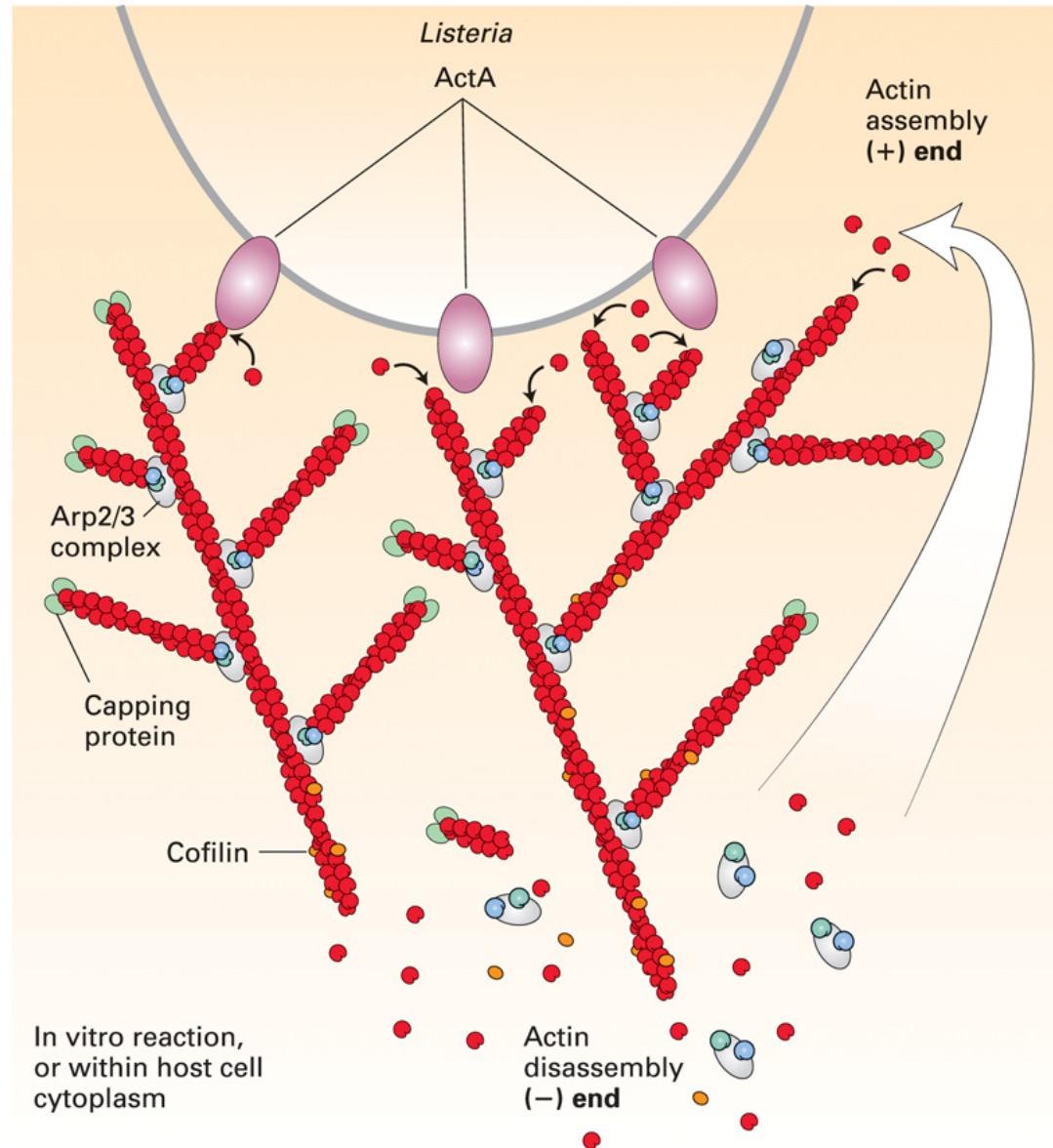
(a)



(b)



(c)





# Actin-based movement powered by myosin - a mechanochemical enzyme.

Common Structure (head, neck and tail) - many variants


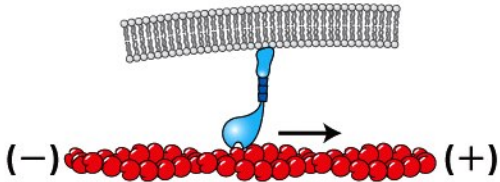
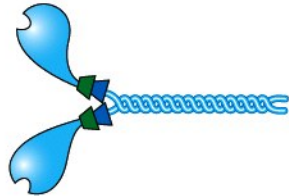
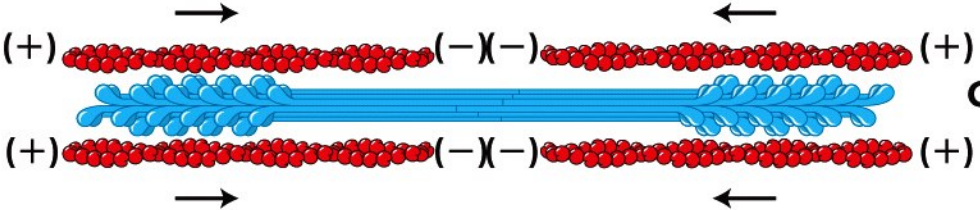
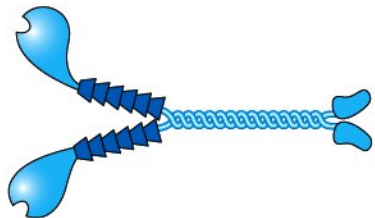
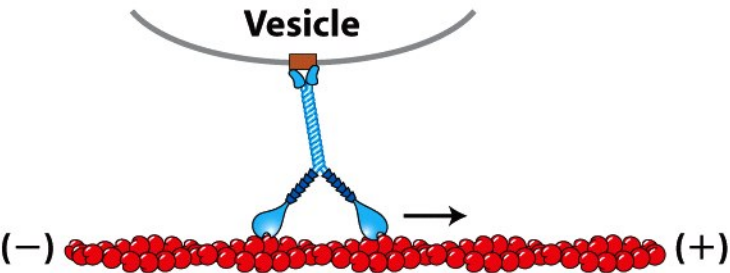
Class	Step size		Function
I 	10–14 nm		Membrane association, endocytosis
II 	5–10 nm		Contraction
V 	36 nm		Organelle transport

Figure 17-23  
Molecular Cell Biology, Sixth Edition  
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# Myosin can move along actin filaments

## Model for the coupling of ATP hydrolysis to movement of myosin along an actin filament

A 'power stroke' is generated through the hydrolysis of ATP (and release of  $P_i$ ) --> moving the myosin along the actin filament

Because myosin heads are tethered by the interaction of tails in a thick filament, the actin filament moves relative to myosin → muscle contraction

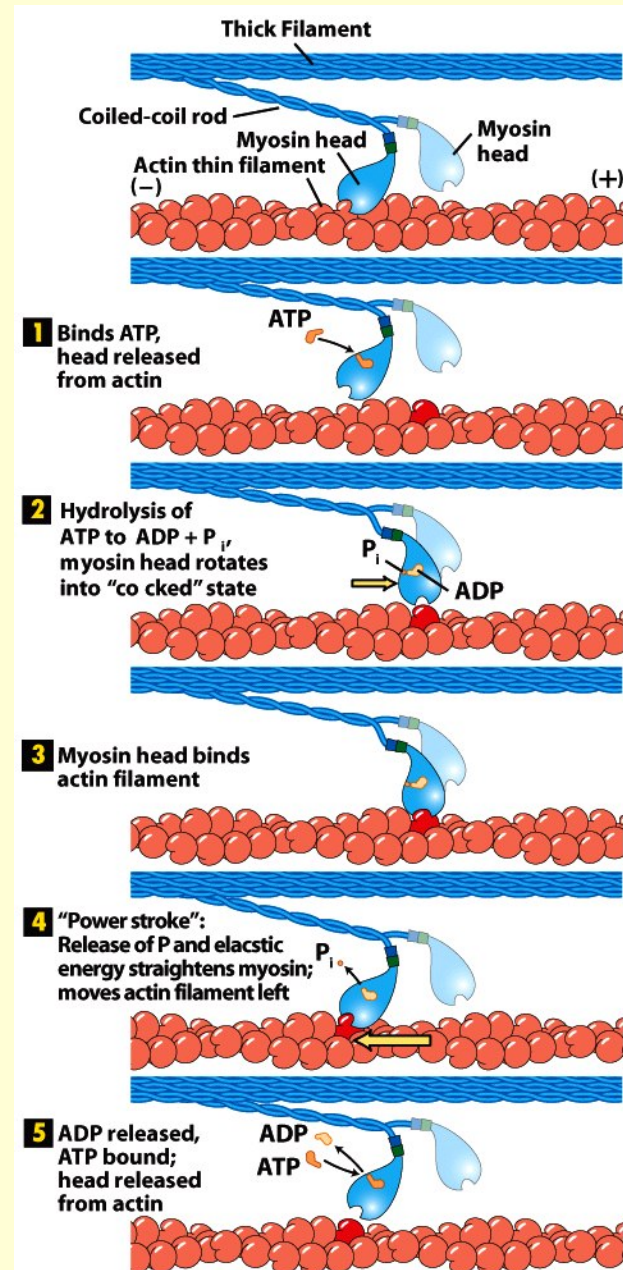
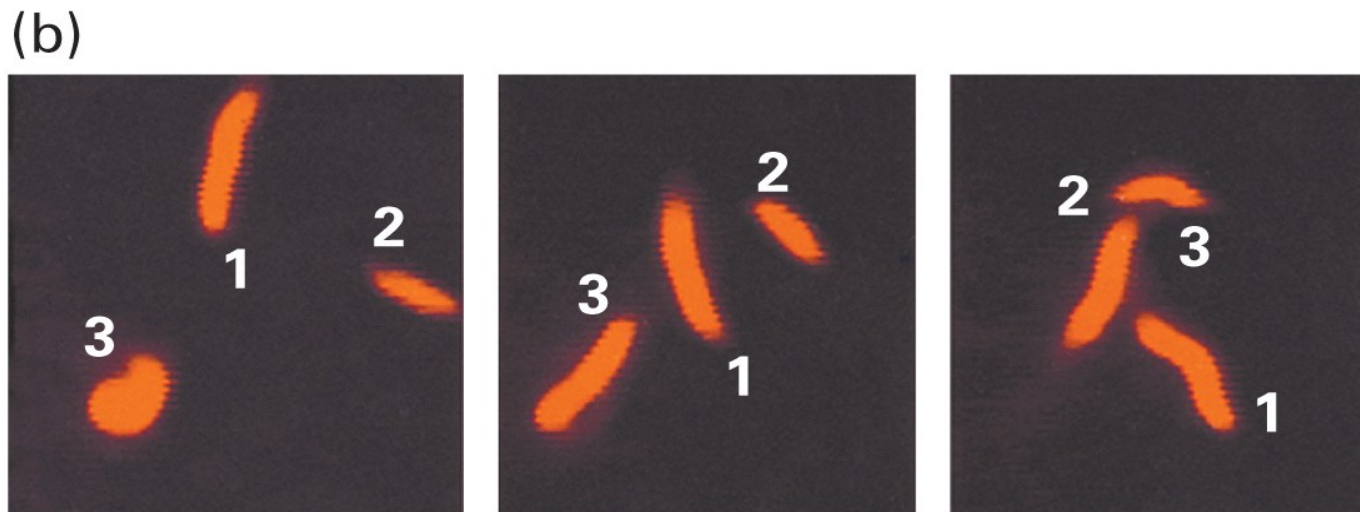
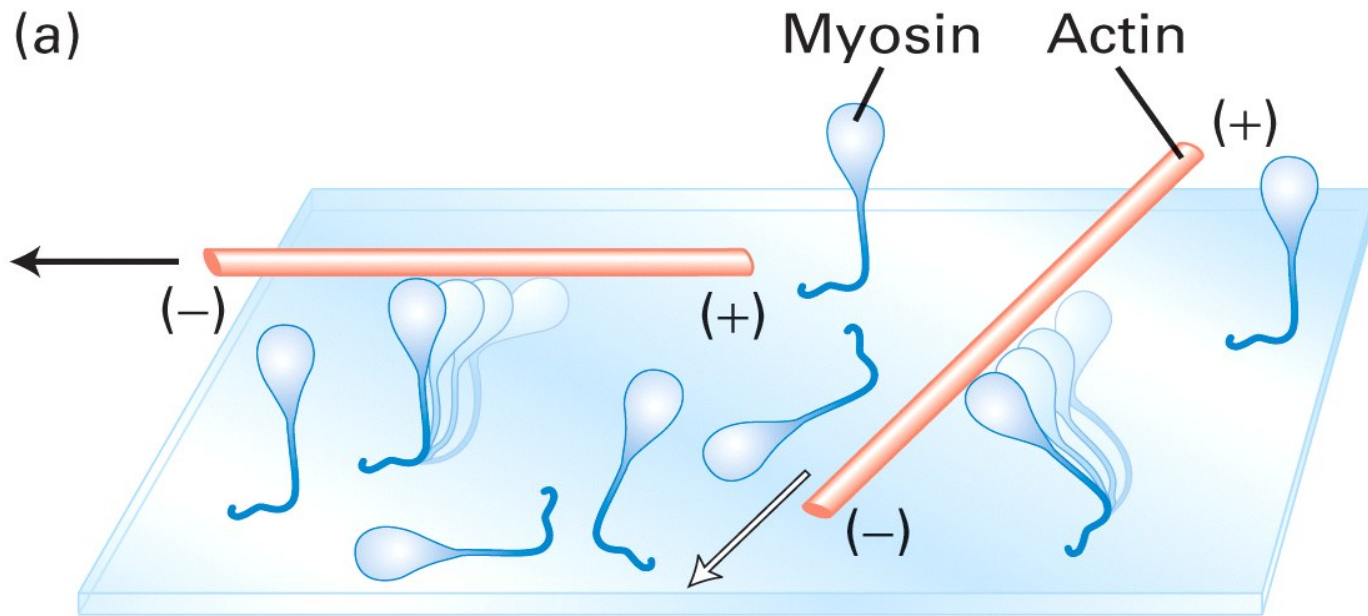


Figure 17-24a  
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Detection of myosin-powered movement of actin filaments by a sliding filament assay (Figure 17-23 Lodish et al., 7th ed)

# The interaction of myosin and actin drive the contraction of skeletal muscle

Muscle cells contain repeating arrays of filament bundles - sarcomeres

Actin filaments (thin filaments)

Myosin filaments (thick filaments -myosin II)

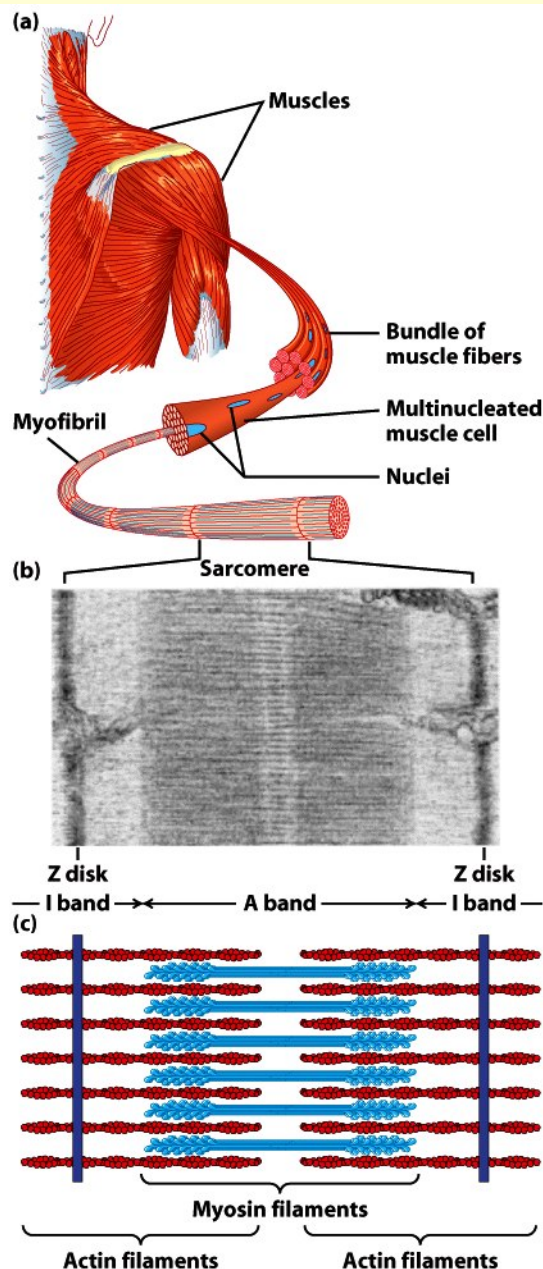


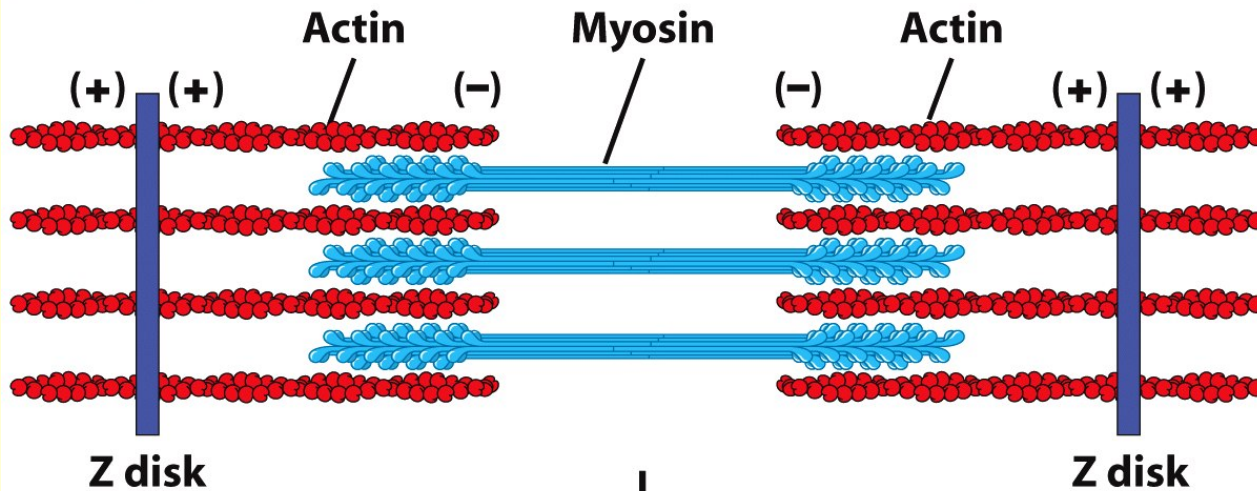
Figure 17-29

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## Relaxed



## Contracted

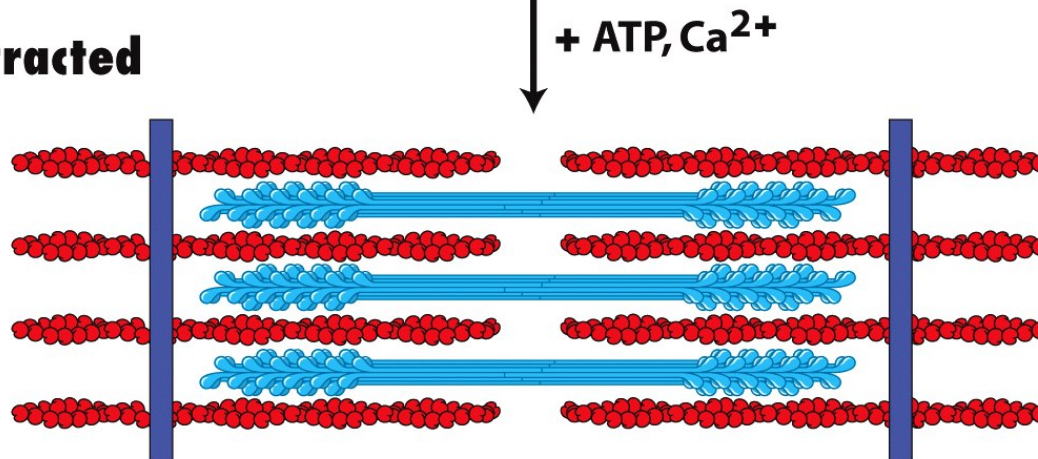


Figure 17-30  
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The *sliding filament* model of muscle contraction; myosin ‘pulls’ actin filaments towards the centre of the sarcomere.



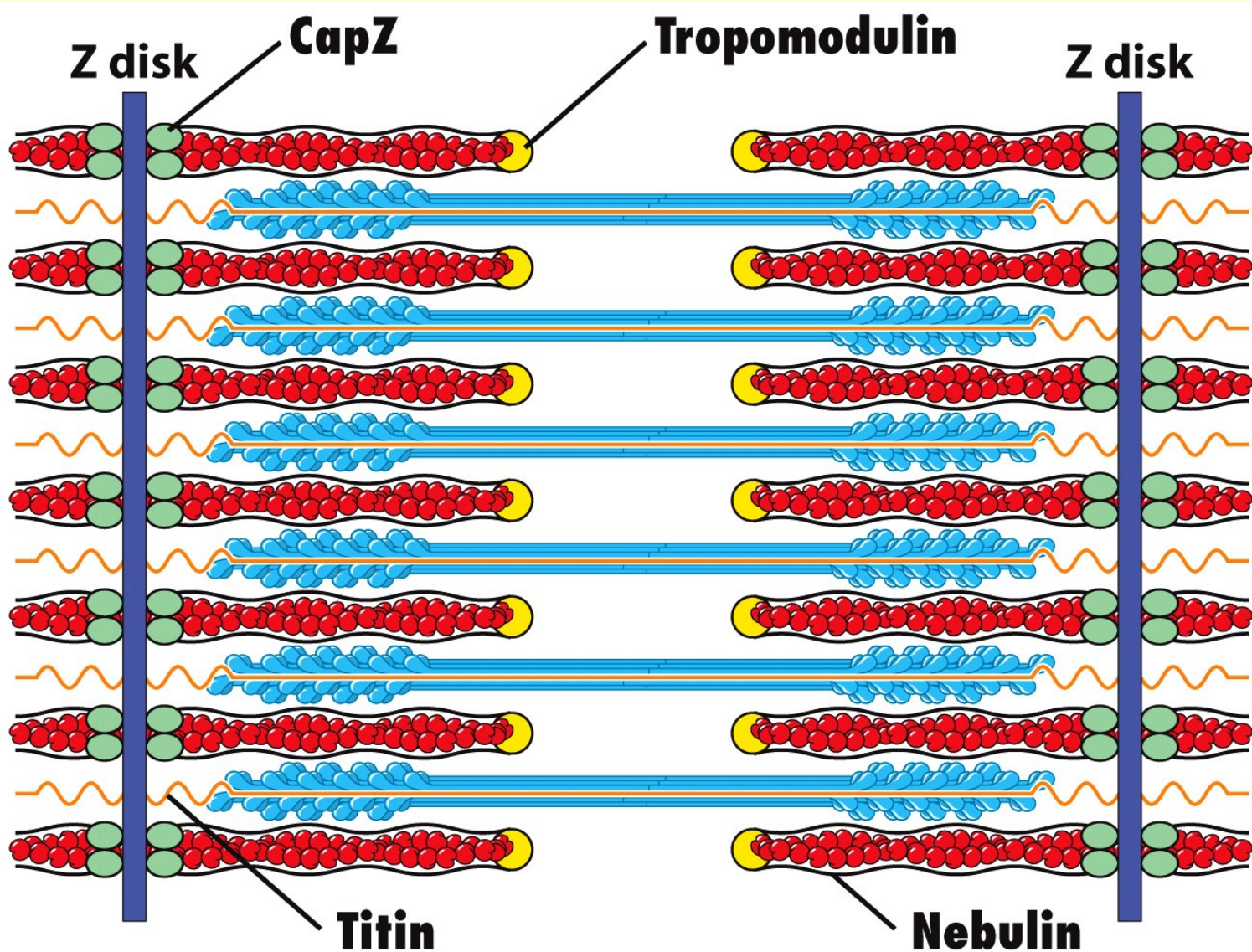


Figure 17-31  
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Accessory proteins maintain the contractile apparatus within each sarcomere

# How is the contraction of muscle regulated ?

Nerve impulse at neuromuscular junction releases  $\text{Ca}^{2+}$

Accessory proteins bound to actin reveal a myosin binding site on the thin filament

Tropomyosin / troponin (TN)

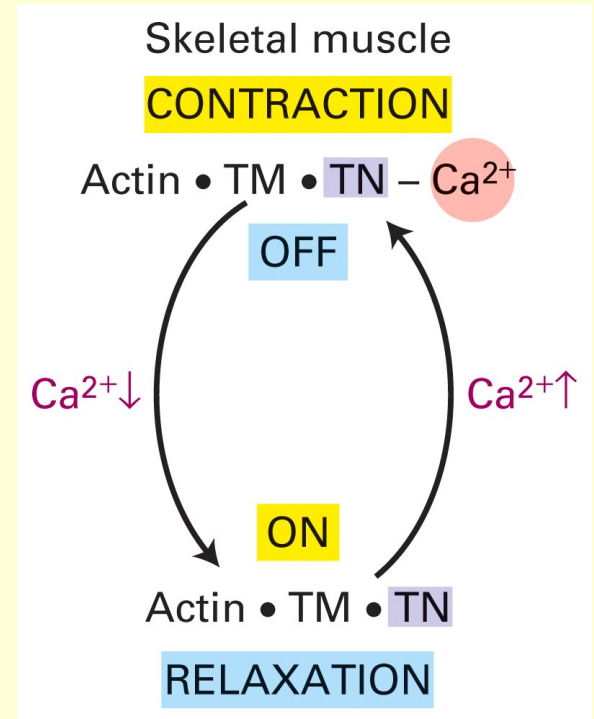
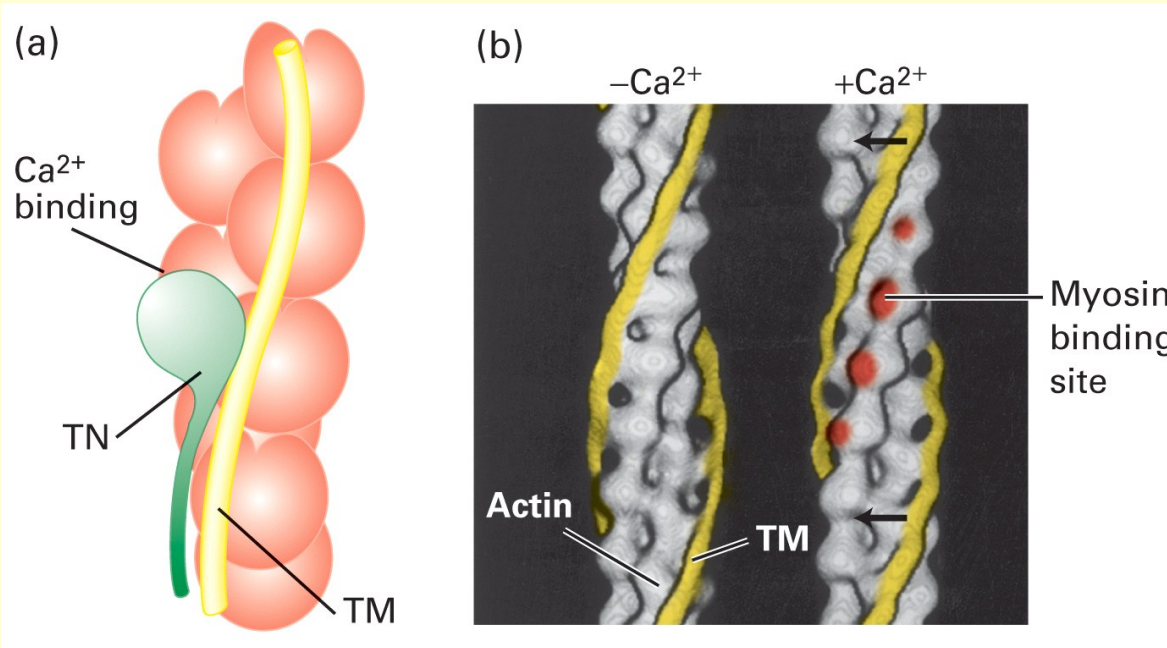
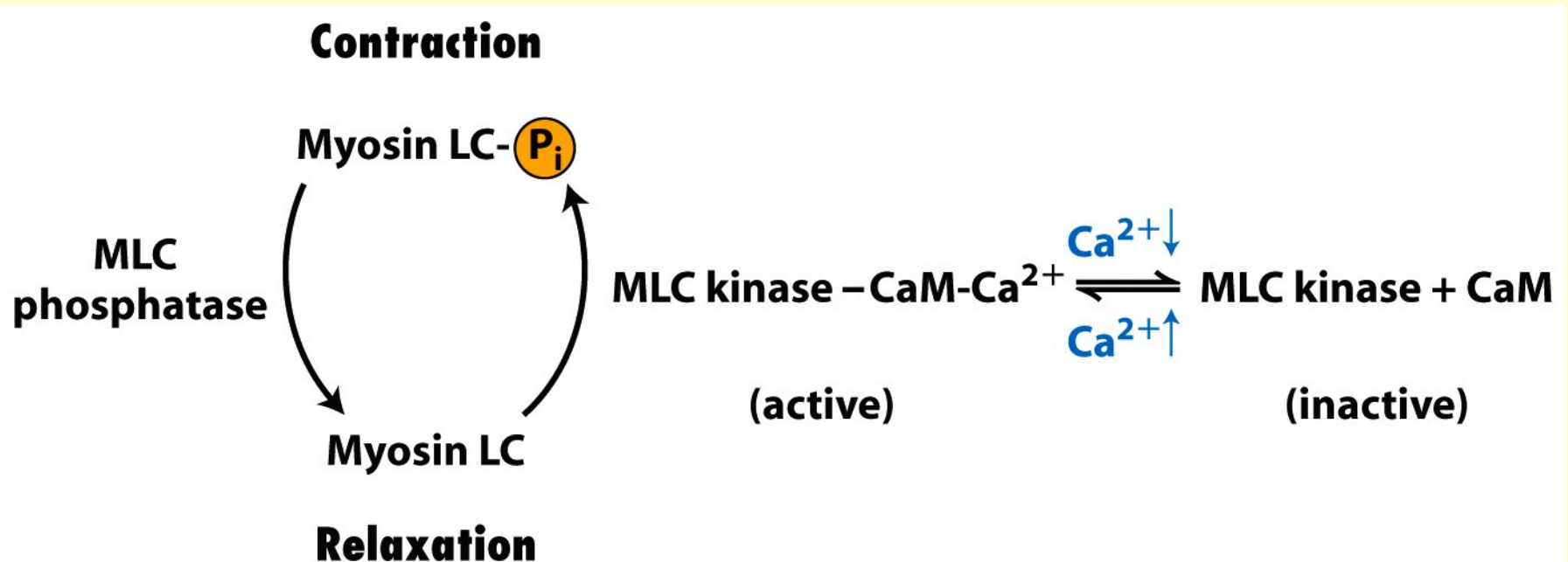


Figure 17-35 Lodish et al., 7th ed

In *smooth* muscle cells, contraction is regulated by the activity of myosin directly *via* phosphorylation of the *myosin regulatory light chain* (LC) by protein kinase and phosphatase enzymes

myosin light chain kinase (MLCK) & myosin light chain phosphatase

Calcium ions ( $\text{Ca}^{2+}$ ) are still the key regulator



## Learning Objectives.

1. Become familiar with the structure of actin and the reversible assembly of F-actin.
2. Understand the role played by actin-binding proteins.
3. Understand how actin is critical to the shape of different types of cells and that cell movement involves actin polymerization.
4. Give examples of how intracellular pathogens use actin polymerization for movement
5. Know the basic features of myosin structure and the function of different classes of myosin in different cell types.
6. Understand how actin and myosin II function together during the contraction of skeletal muscle and know how this process is regulated by other actin-binding proteins.