

Essential Cell Biology

Third Edition

Chapter 4

Protein Structure and Function

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2生化 岡島 H30.05.14

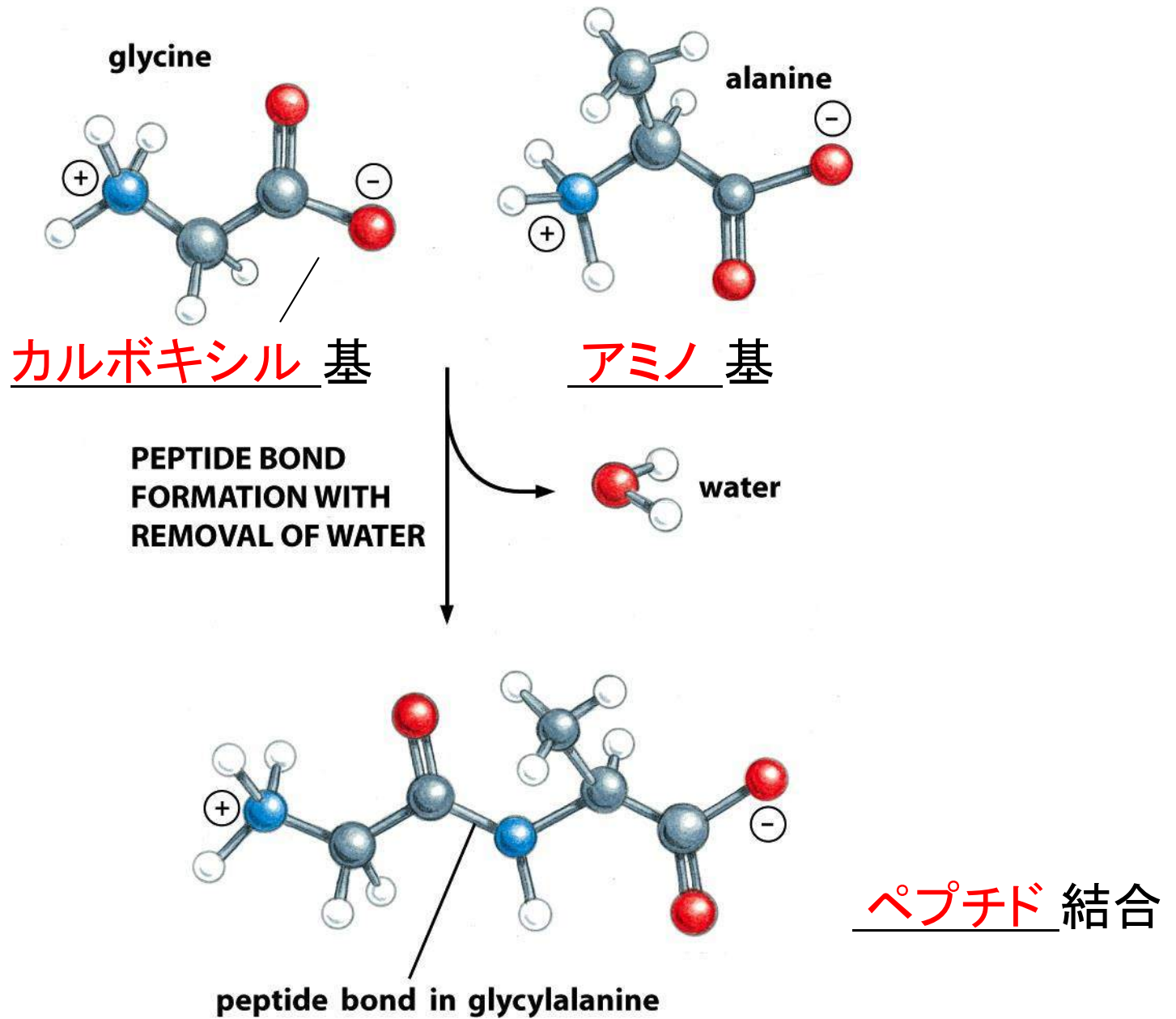


Figure 4-1 *Essential Cell Biology*, 3rd ed. (© Garland Science 2010) 121p.

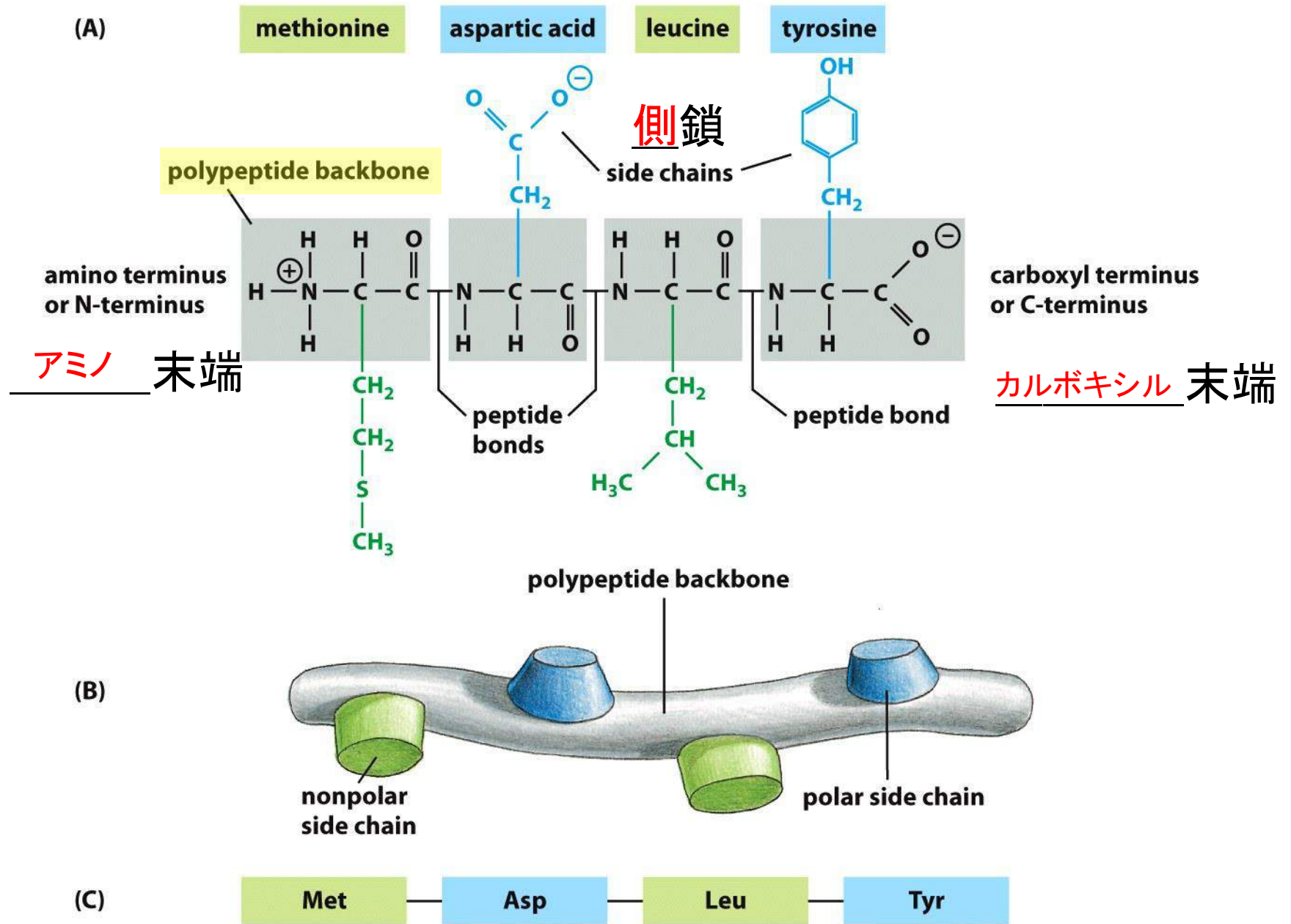


Figure 4-2 *Essential Cell Biology*, 3rd ed. (© Garland Science 2010) 122p.

側鎖の化学的性質によるアミノ酸の分類

AMINO ACID		SIDE CHAIN	
Aspartic acid	Asp	D	negative
Glutamic acid	Glu	E	negative
Arginine	Arg	R	positive
Lysine	Lys	K	positive
Histidine	His	H	positive
Asparagine	Asn	N	uncharged polar
Glutamine	Gln	Q	uncharged polar
Serine	Ser	S	uncharged polar
Threonine	Thr	T	uncharged polar
Tyrosine	Tyr	Y	uncharged polar

AMINO ACID		SIDE CHAIN	
Alanine	Ala	A	nonpolar
Glycine	Gly	G	nonpolar
Valine	Val	V	nonpolar
Leucine	Leu	L	nonpolar
Isoleucine	Ile	I	nonpolar
Proline	Pro	P	nonpolar
Phenylalanine	Phe	F	nonpolar
Methionine	Met	M	nonpolar
Tryptophan	Trp	W	nonpolar
Cysteine	Cys	C	nonpolar

POLAR AMINO ACIDS

(hydrophilic)

親水性(極性) アミノ酸

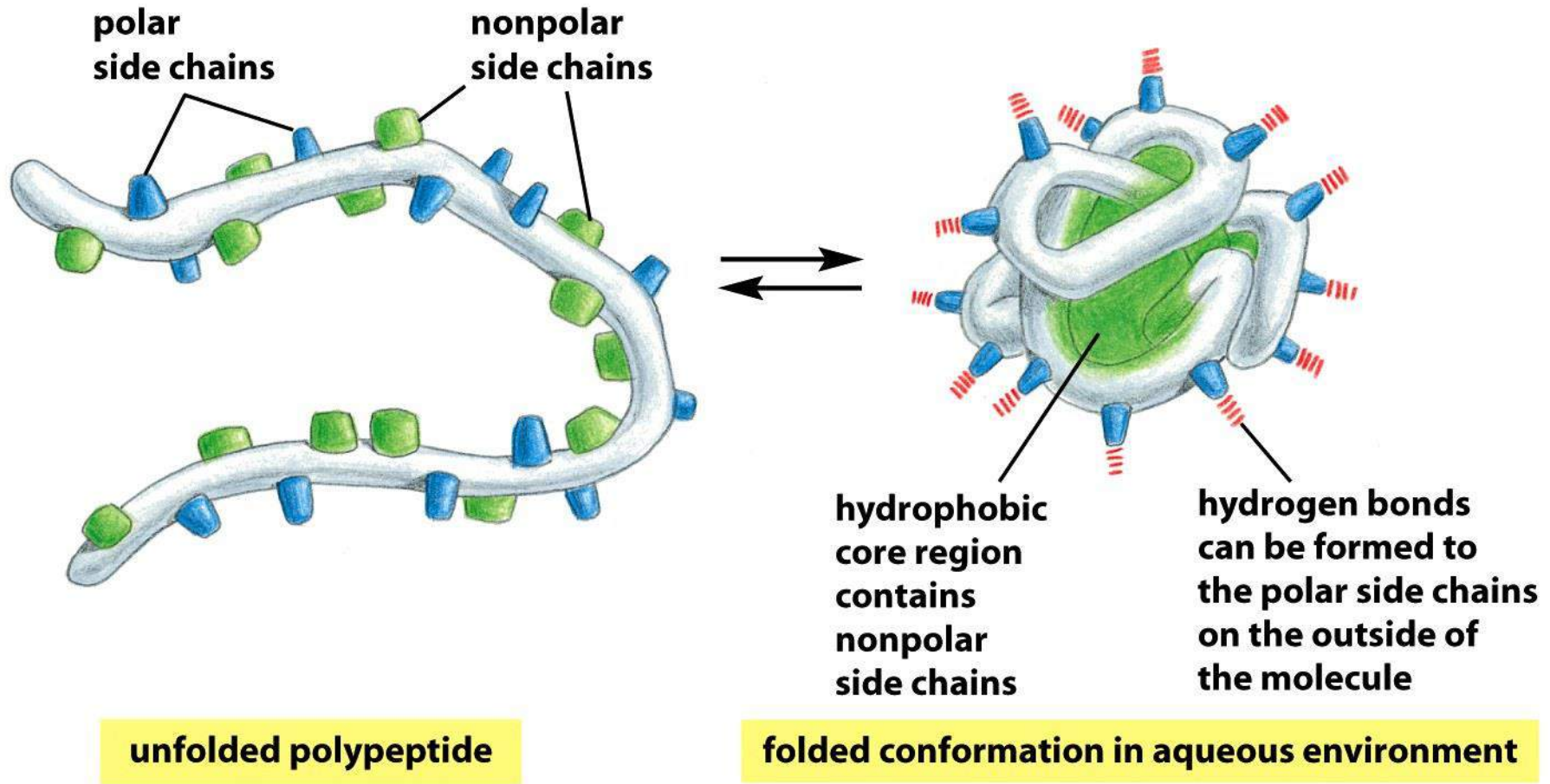
NONPOLAR AMINO ACIDS

(hydrophobic)

疎水性(非極性) アミノ酸

Proteins fold into a conformation of lowest energy

(水分子と水素結合)



Three types of noncovalent bonds help proteins fold

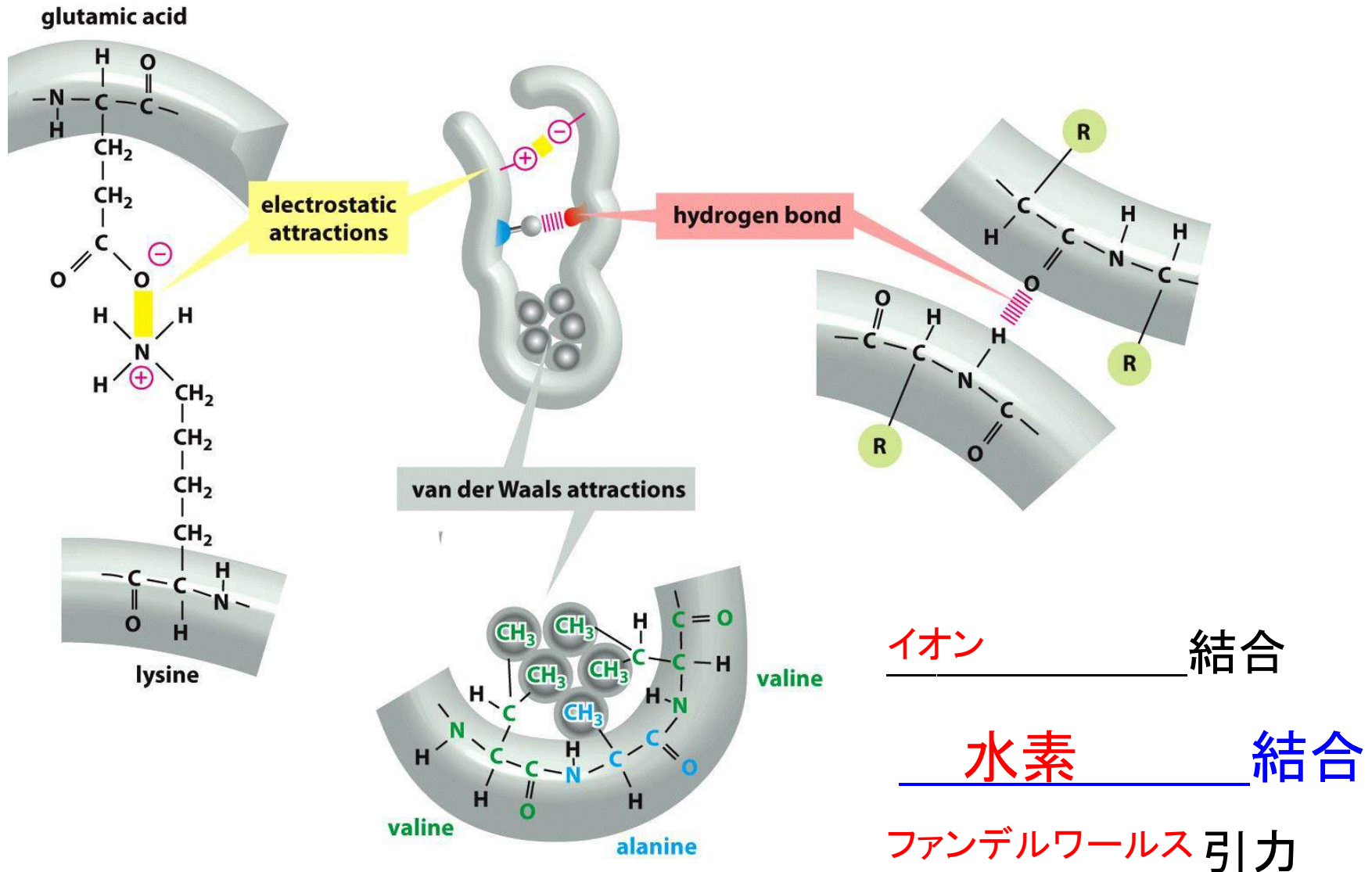
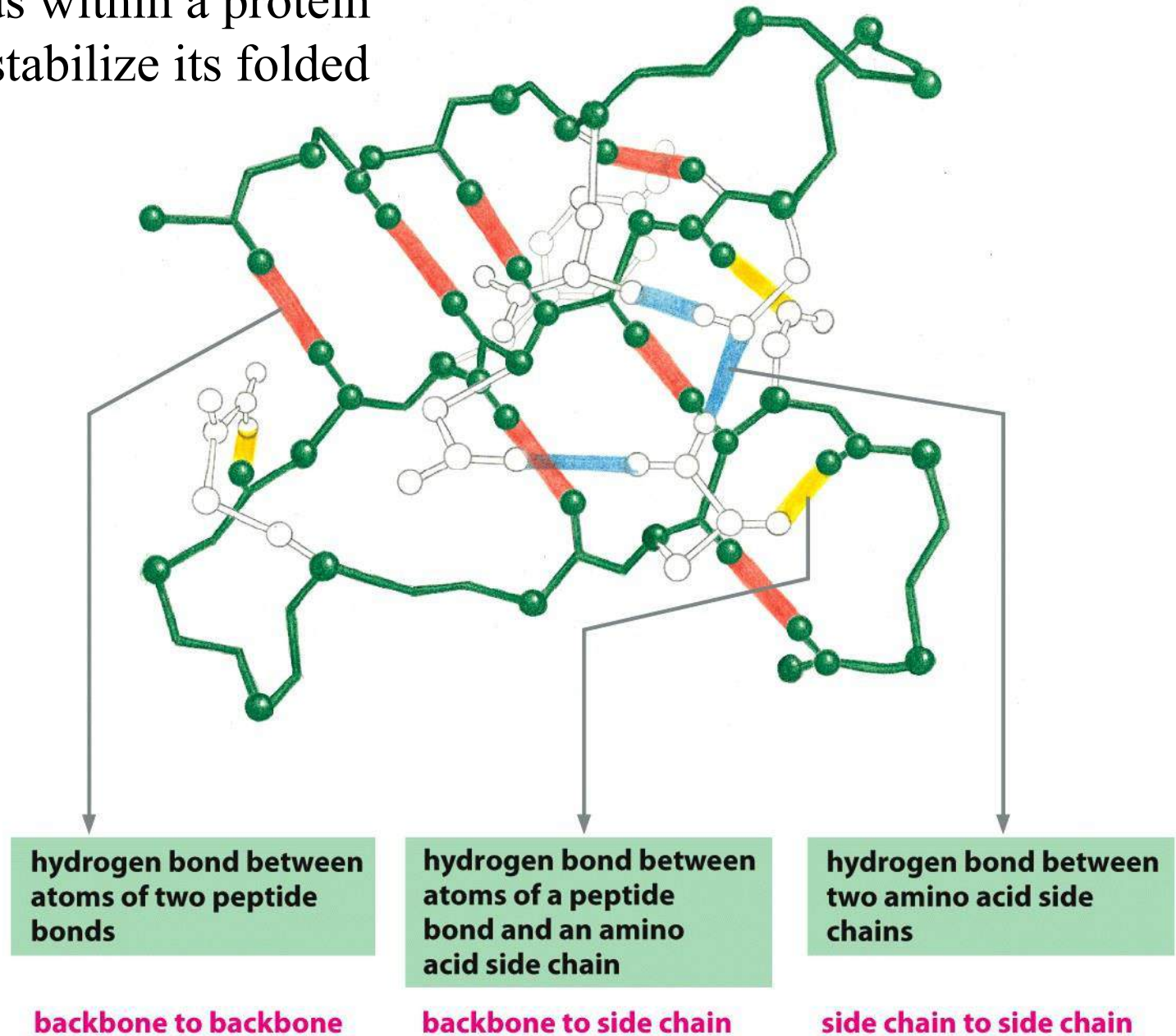


Figure 4-4 *Essential Cell Biology*, 3rd ed. (© Garland Science 2010)123p.

Hydrogen bonds within a protein molecule help stabilize its folded shape



Aggregated Proteins

Neurodegenerative disorders

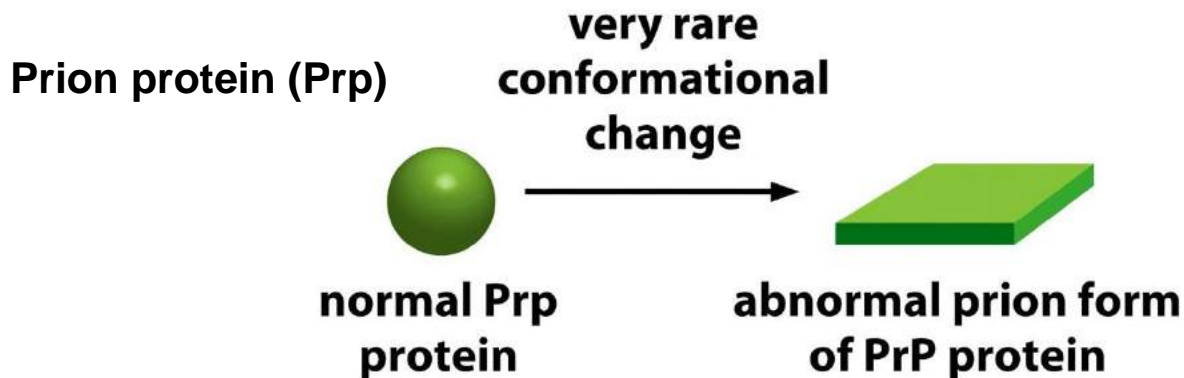
Alzheimer's disease

Huntington's disease

Prion disease

- scrapie---sheep
- Bovine spongiform encephalopathy (BSE)
- Creutzfeldt-Jacob disease (CJD)

prion protein can adopt an abnormal, misfolded form



misfolded protein can induce formation of protein aggregates

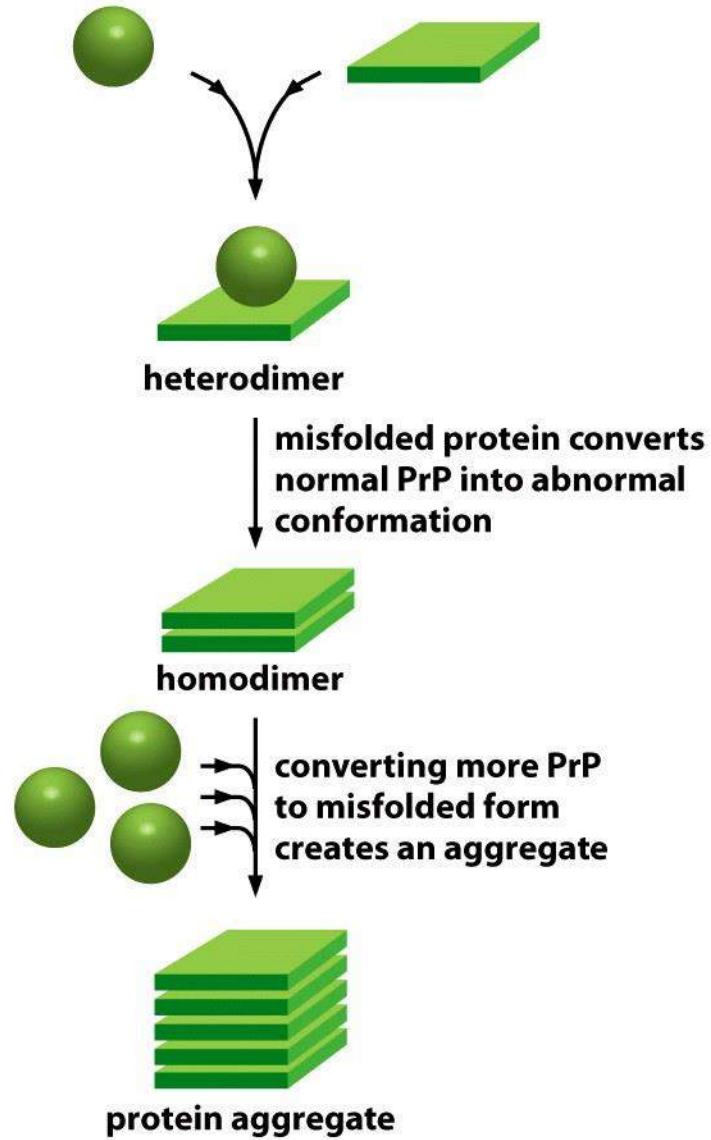


Figure 4-8(B) *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 125p.

Proteins come in a wide variety of complicated shape

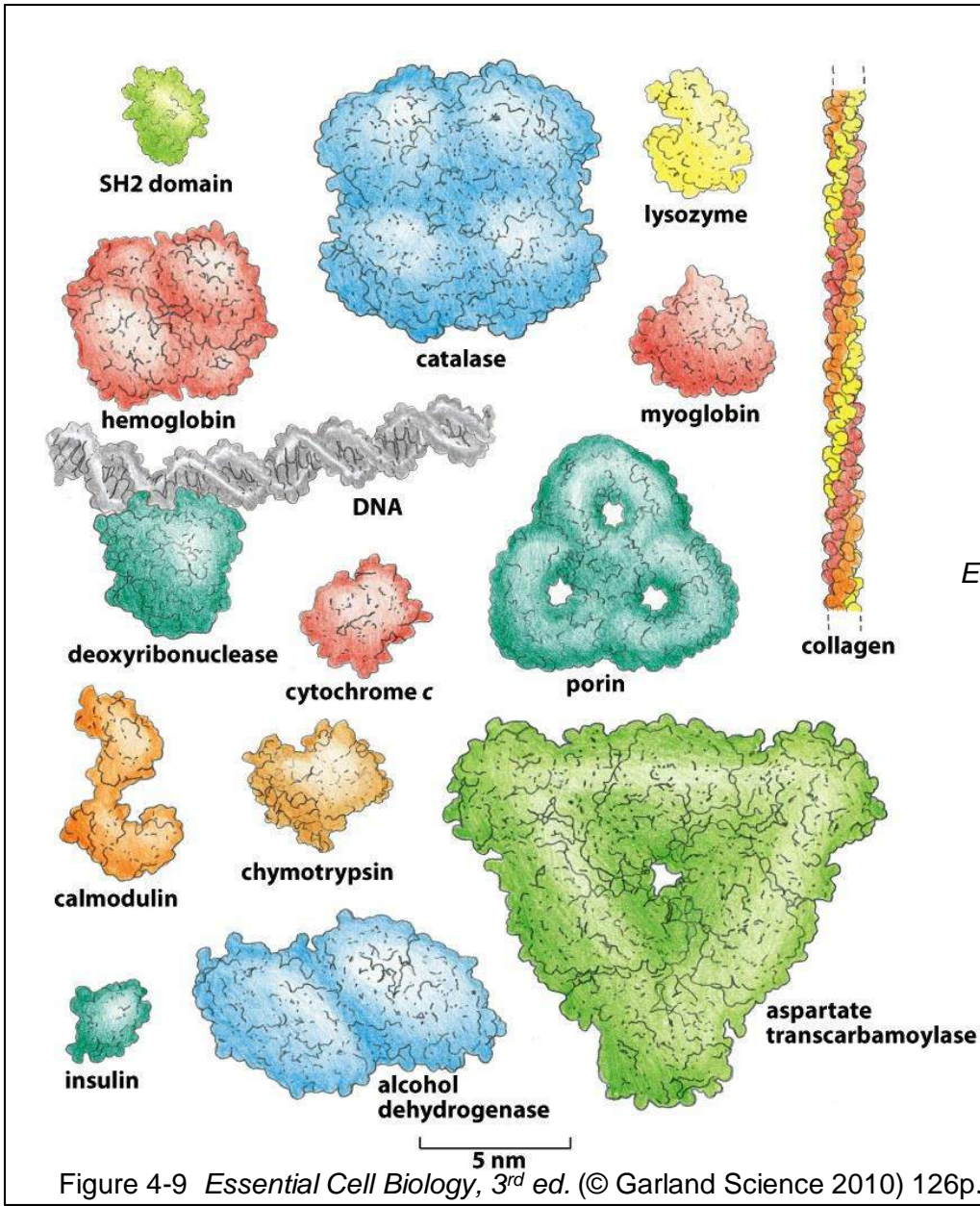
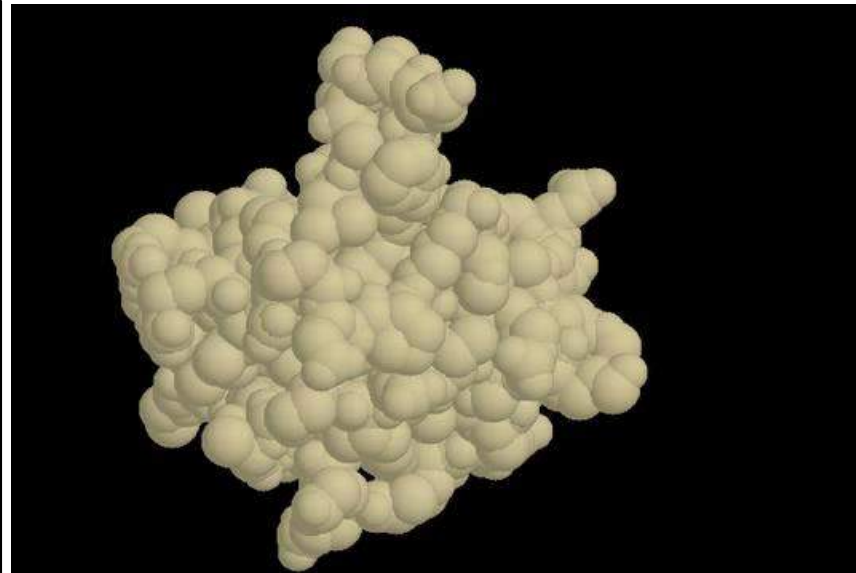


Figure 4-9 *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 126p.



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SH2 domain

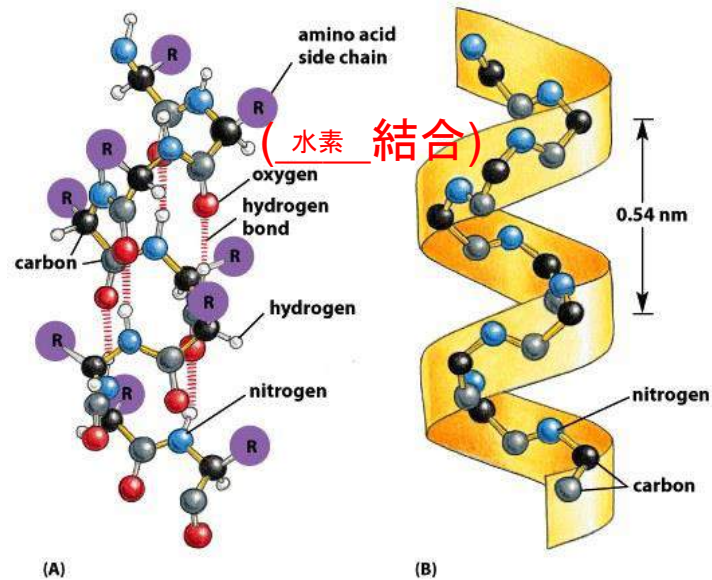
(リン酸化されたチロシンに結合)

polypeptide backbone model
 ribbon model

Wire model
 Space-filling model } side chain

Tertiary structure

(三次構造)



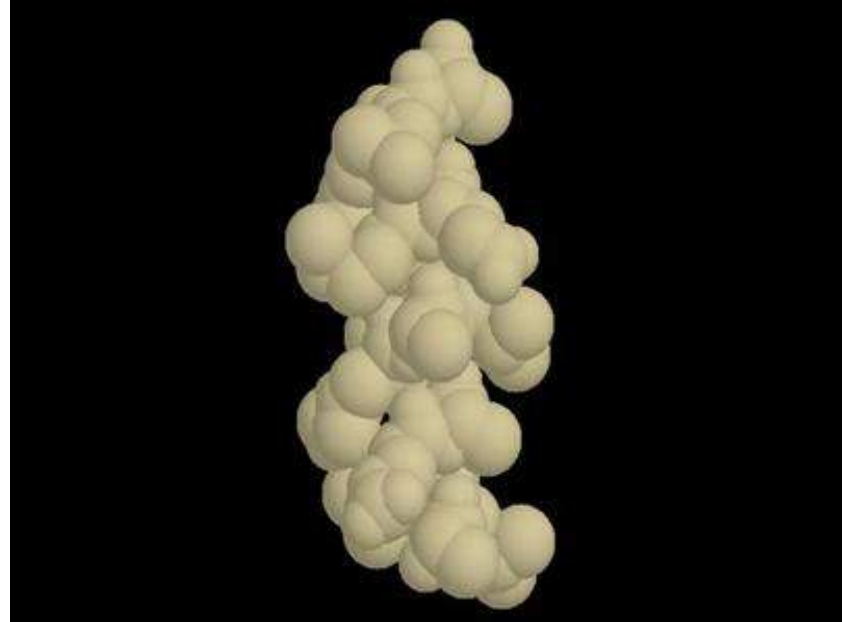
(A)

(B)

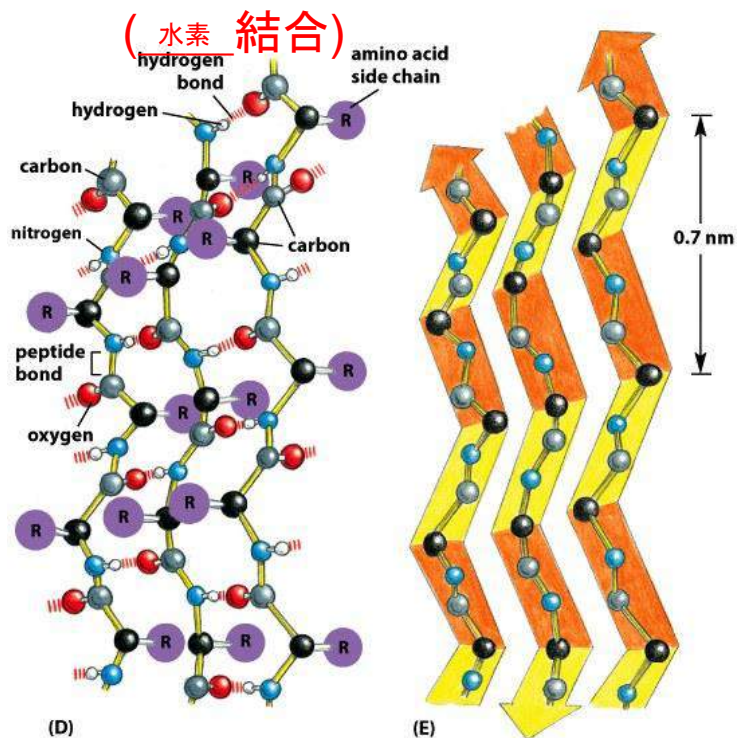
α helix



(C)



水素結合の位置は?
側鎖の位置は?



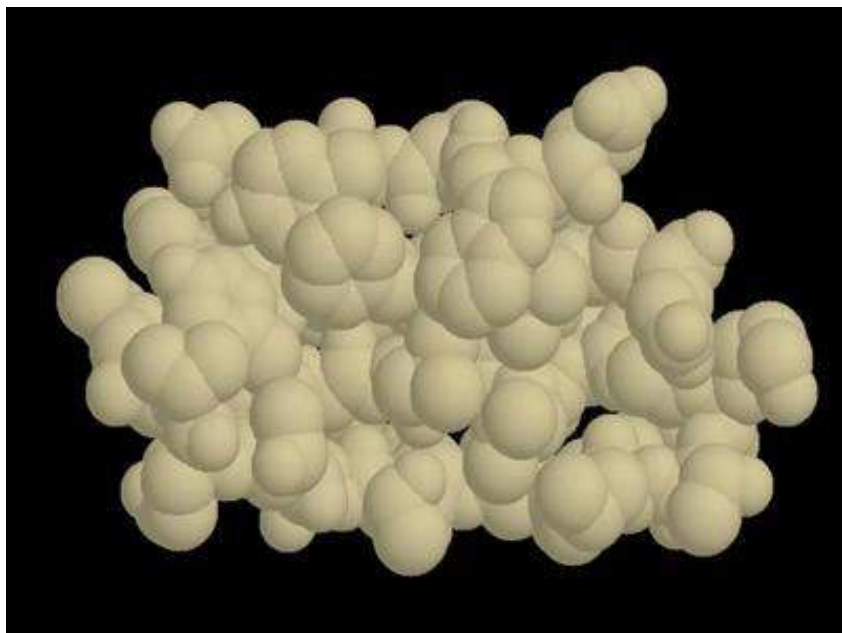
(D)

(E)

β sheet



(F)



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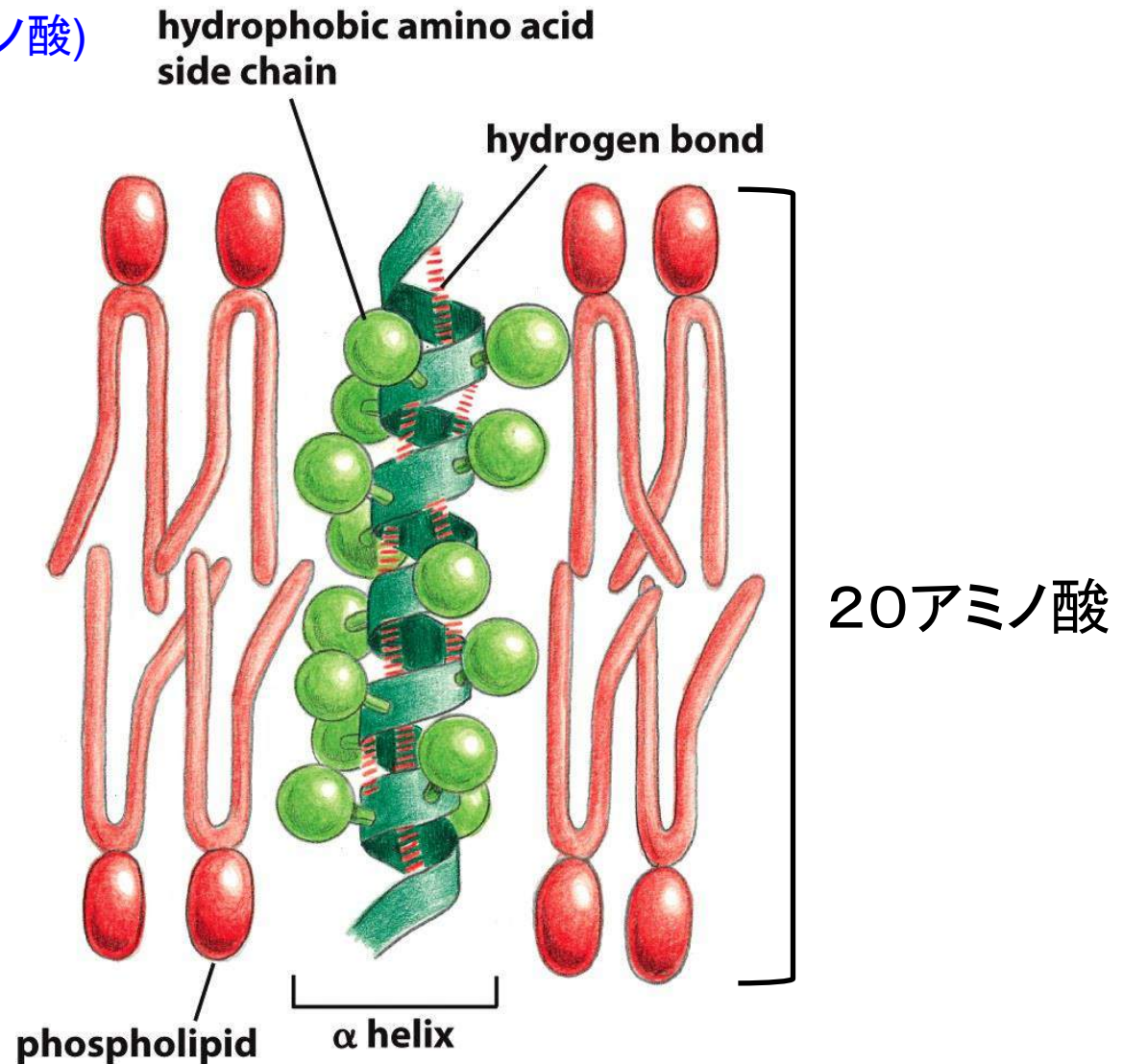
two-dimensional structure

(2次構造)

Figure 4-10 *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 130p.

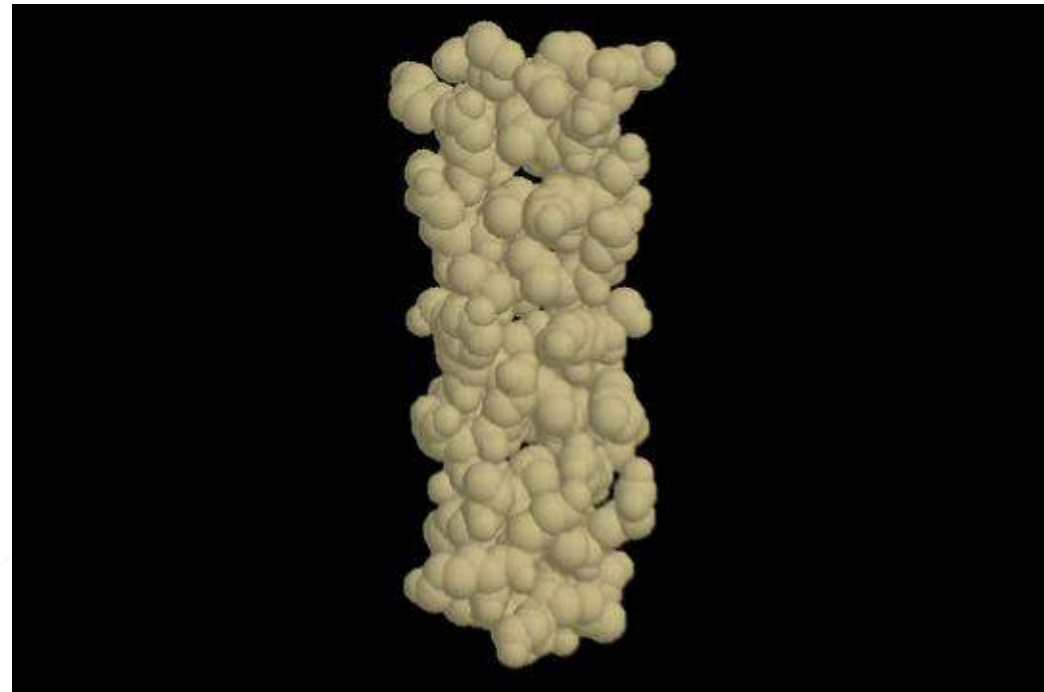
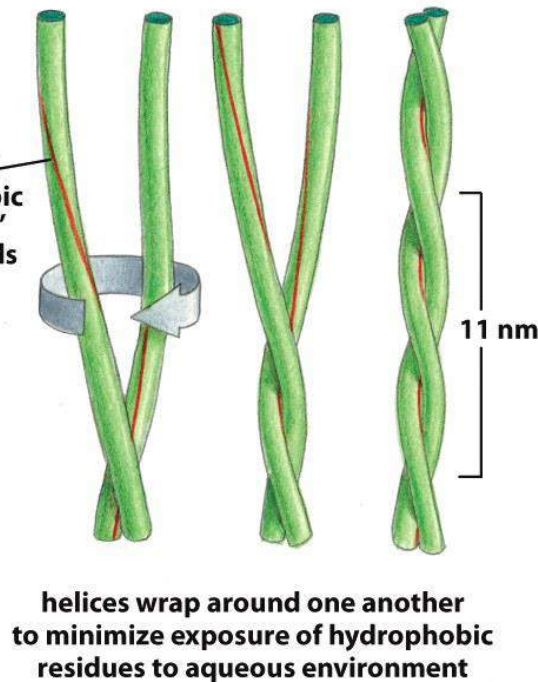
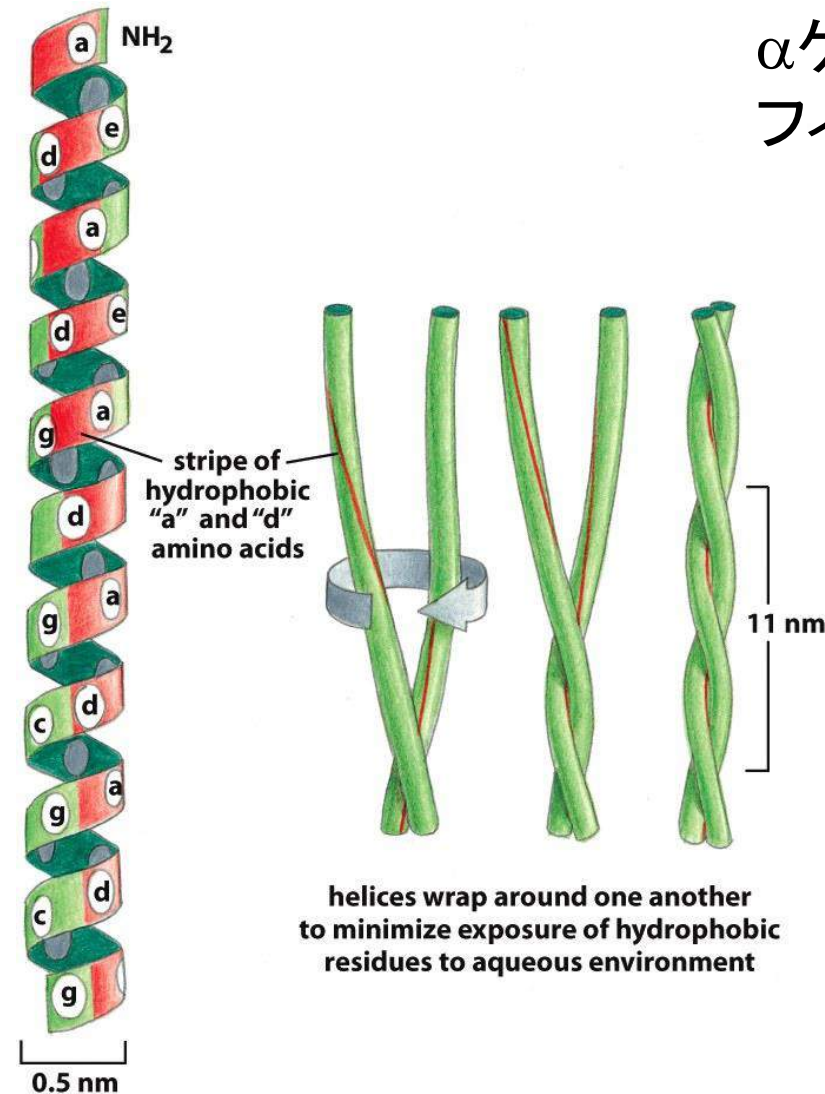
A segment of α -helix can cross a lipid bilayer

20程度の(疎水性アミノ酸)



Two or three α -helices form **coiled-coil**

α ケラチン、ミオシンの2重らせん構造
フィブリノーゲンの3重らせん構造

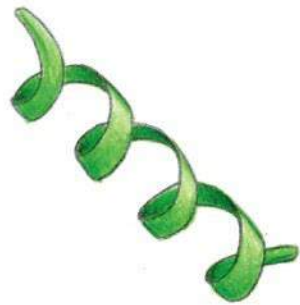


HOOC-COOH
Essential Cell Biology, 3rd ed. (© Garland Science 2010) 付属DVDより

(A) (B) (C)

Figure 4-13 *Essential Cell Biology*, 3rd ed. (© Garland Science 2010) 132p.

Many proteins are composed of separate functional domains
(ドメイン)

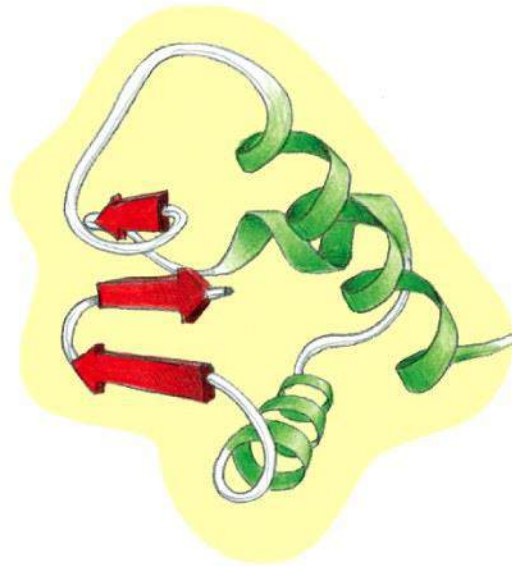


α helix

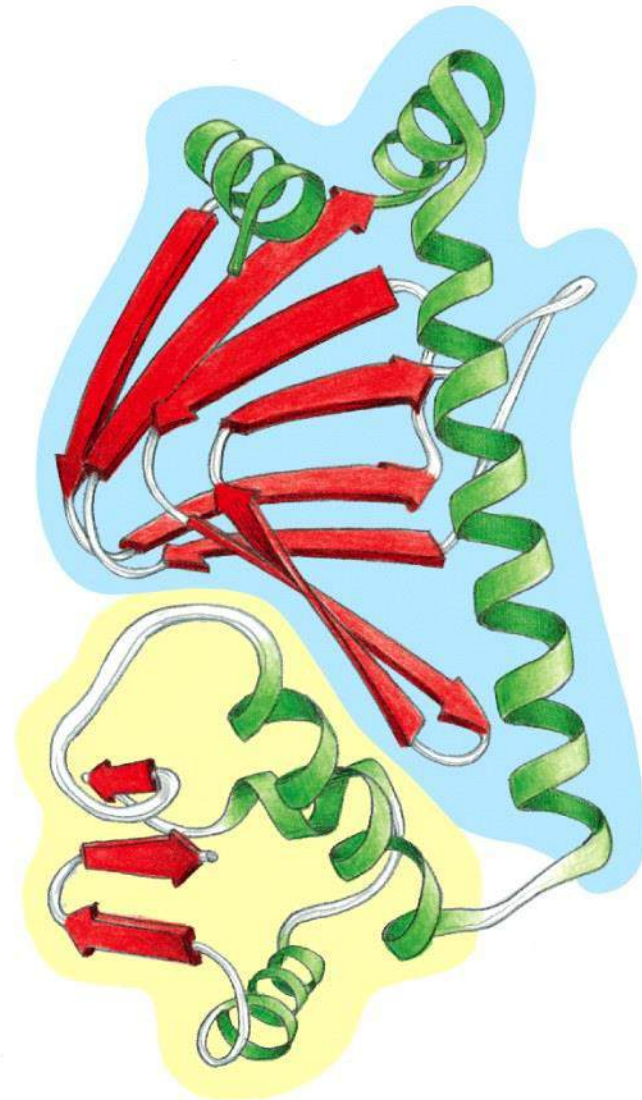


β sheet

secondary structure

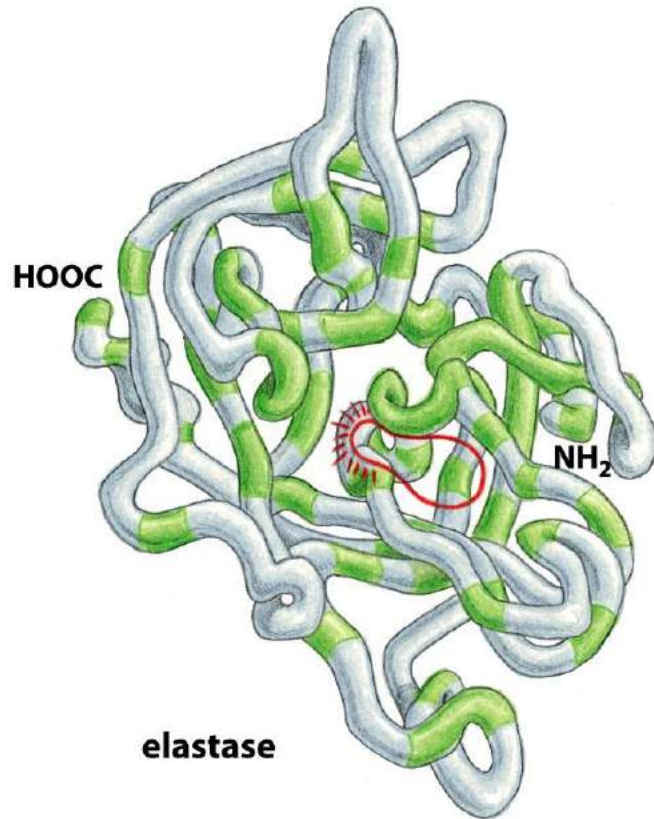


single polypeptide domain

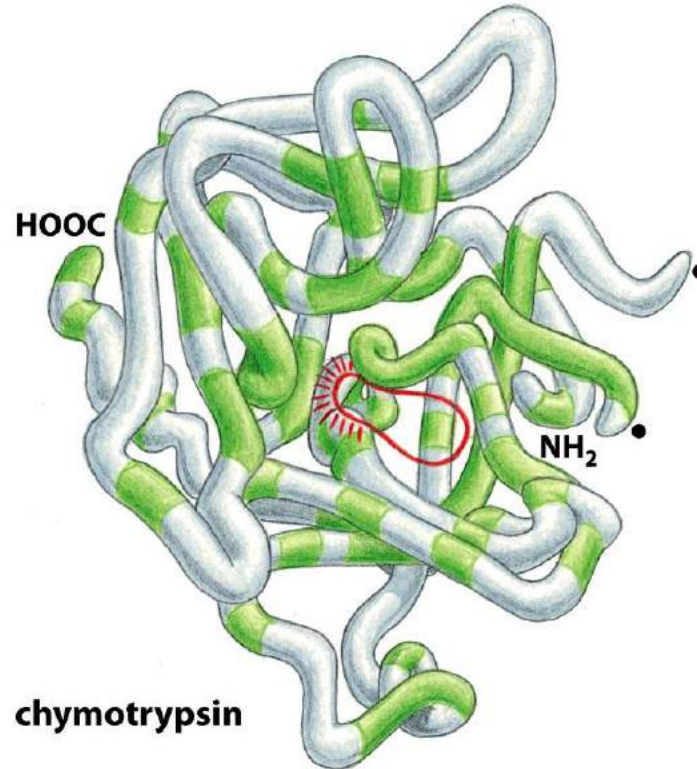


protein molecule made of two different domains

Serine proteases comprise a family of proteolytic enzymes



エラスチンを分解

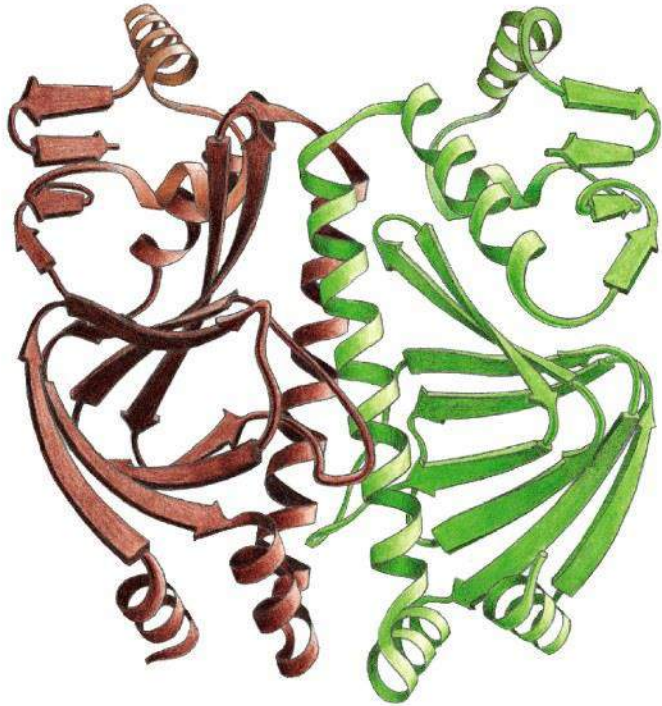


芳香族アミノ酸の
カルボキシル基側を加水分解

タンパク質ファミリー
アミノ酸配列の類似性
構造の類似性

Many proteins molecules contain multiple copies of a single protein subunit

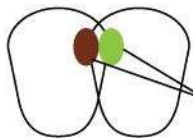
(サブユニット)



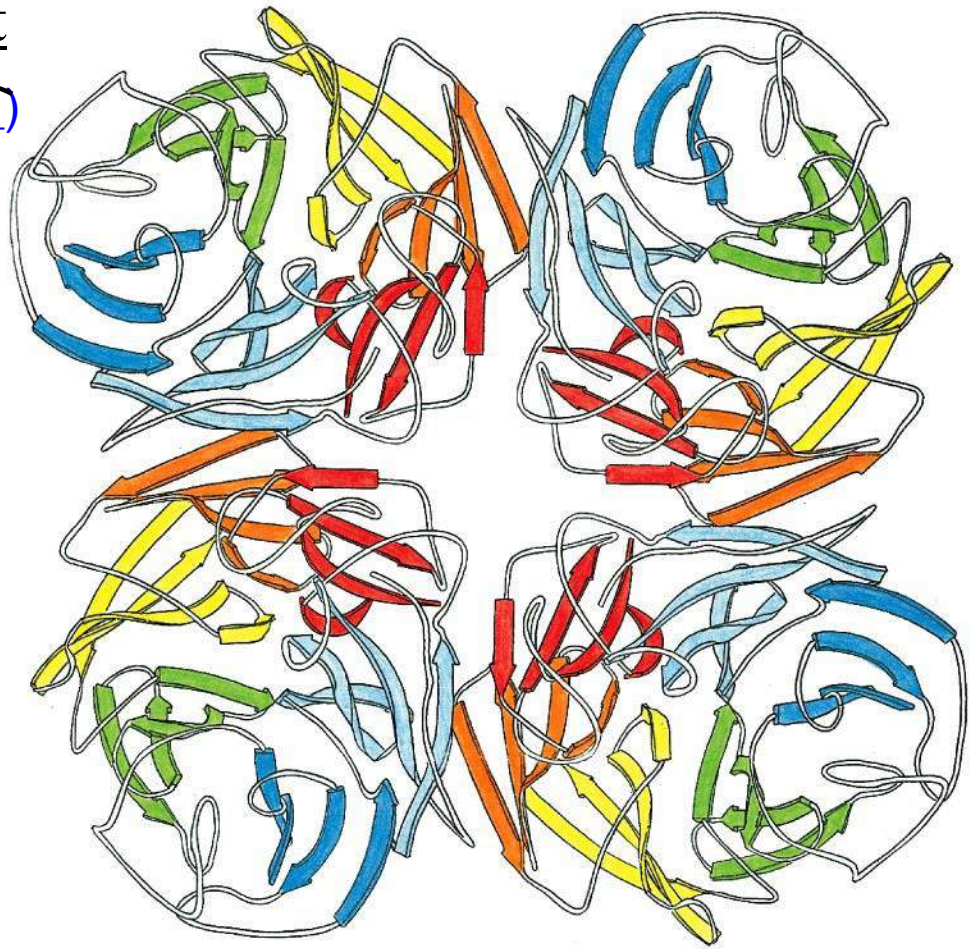
dimer of the CAP protein

2量体

(A)



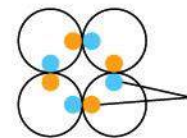
identical binding site on each monomer



tetramer of neuraminidase protein

4量体

(B)



two non-identical binding sites on each monomer

ヘモグロビン

4量体(ホモ2量体のヘテロ2量体)

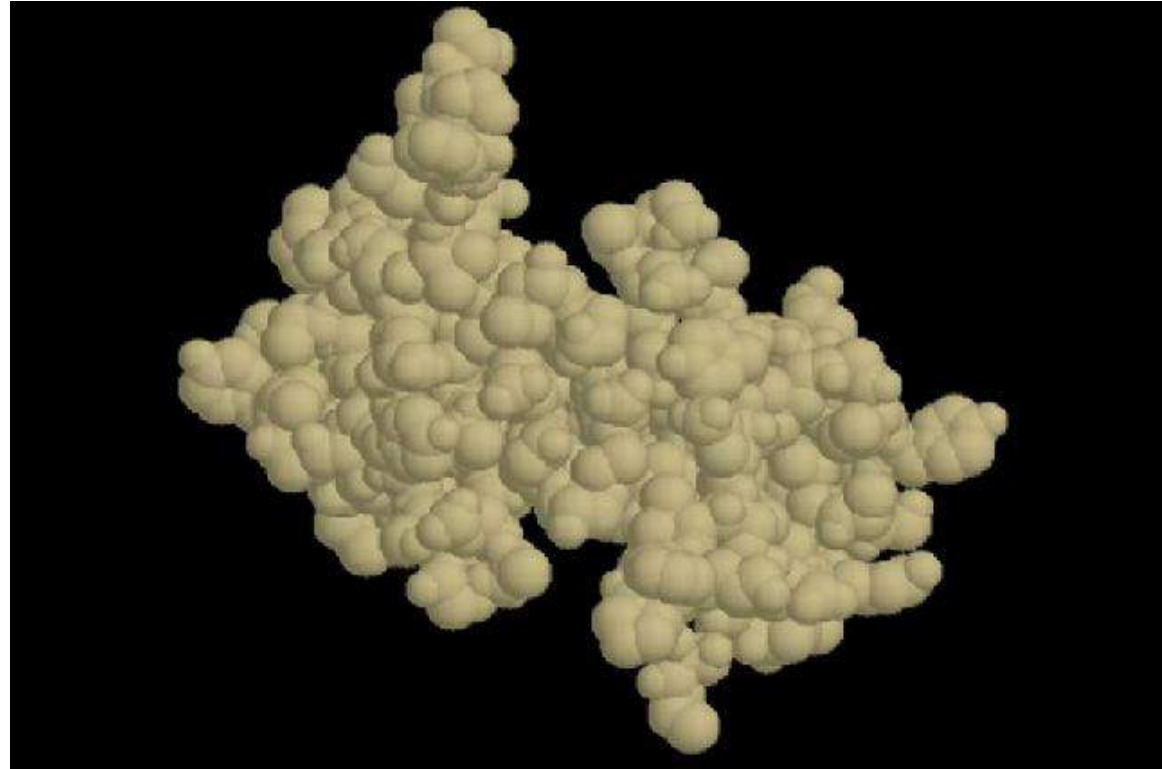
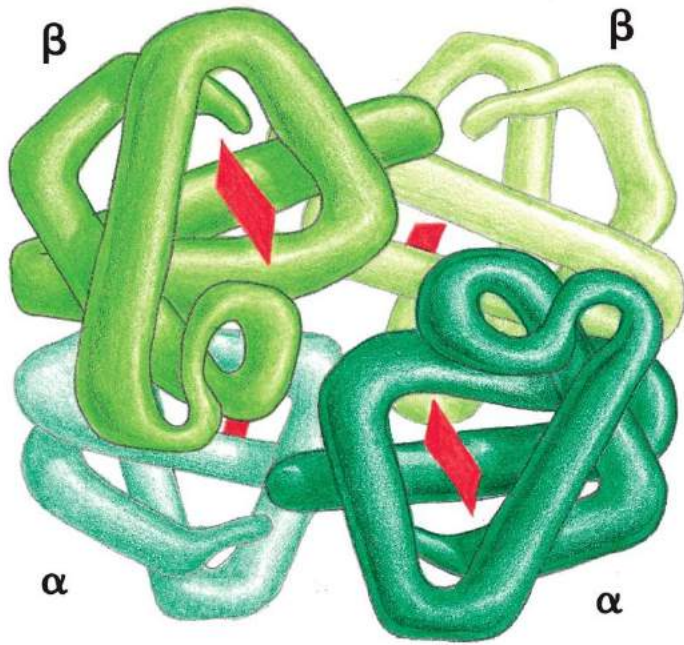


Figure 4-20 *Essential Cell Biology, 3rd ed.*
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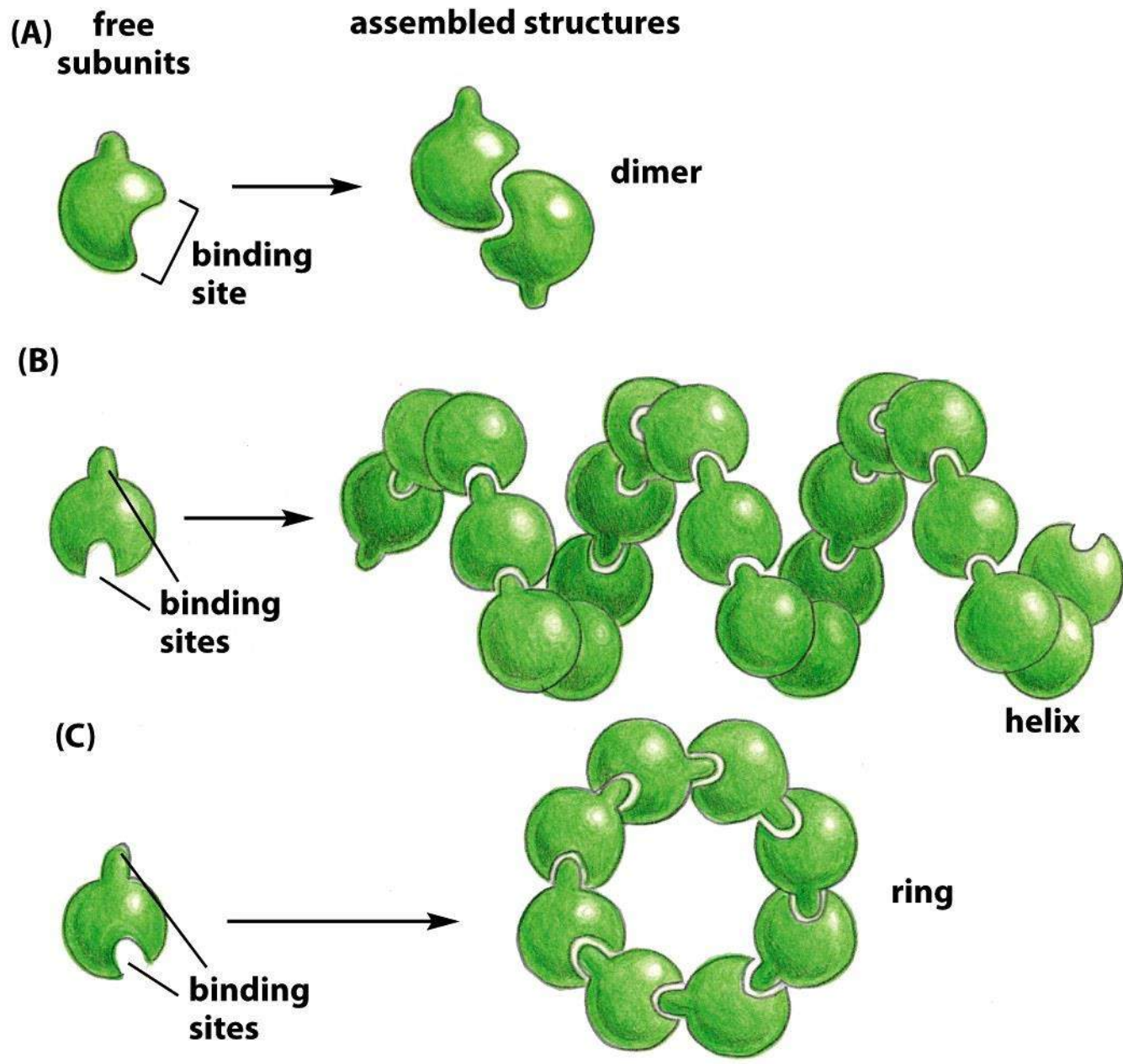


Figure 4-21 *Essential Cell Biology*, 3rd ed. (© Garland Science 2010) 137p.

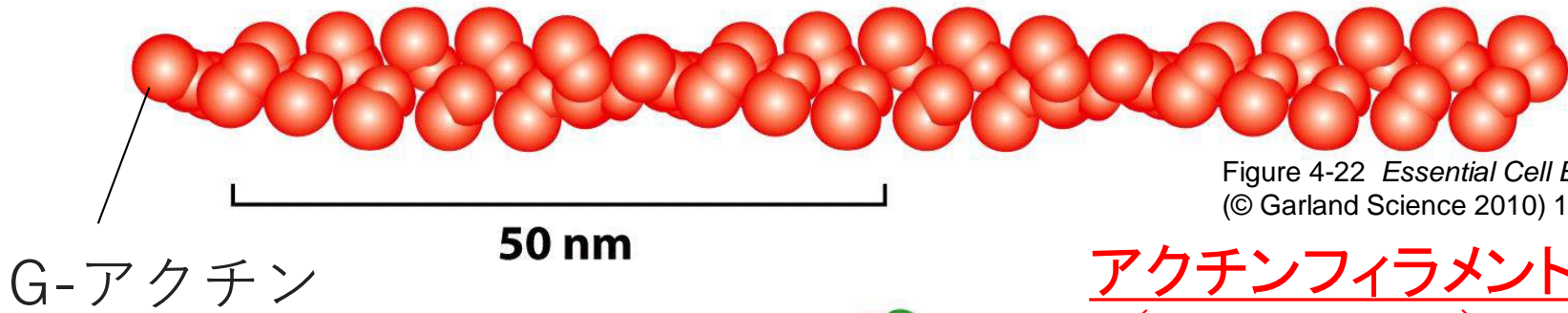


Figure 4-22 *Essential Cell Biology, 3rd ed.*
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アクチンフィラメント
 (F-アクチン)

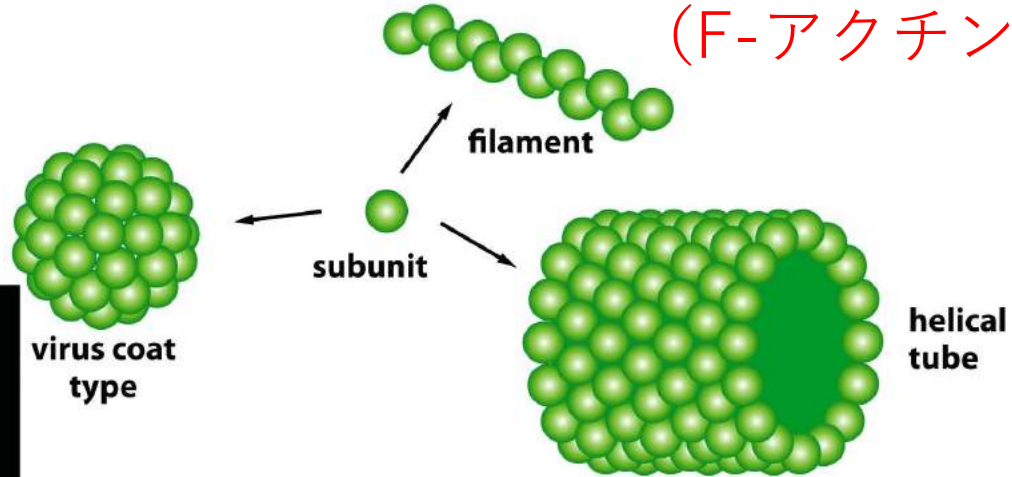


Figure 4-23 *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 138p.

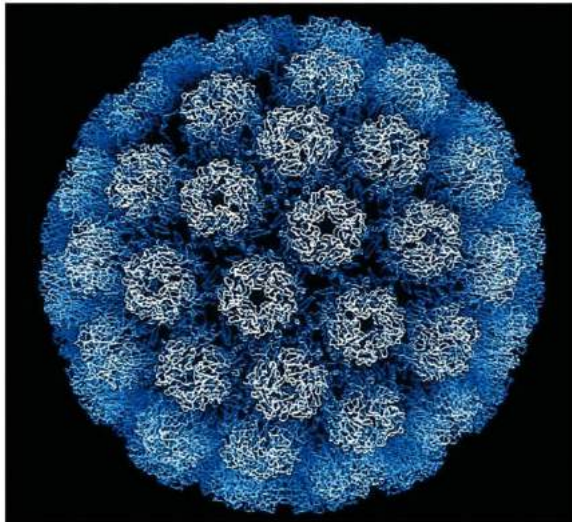


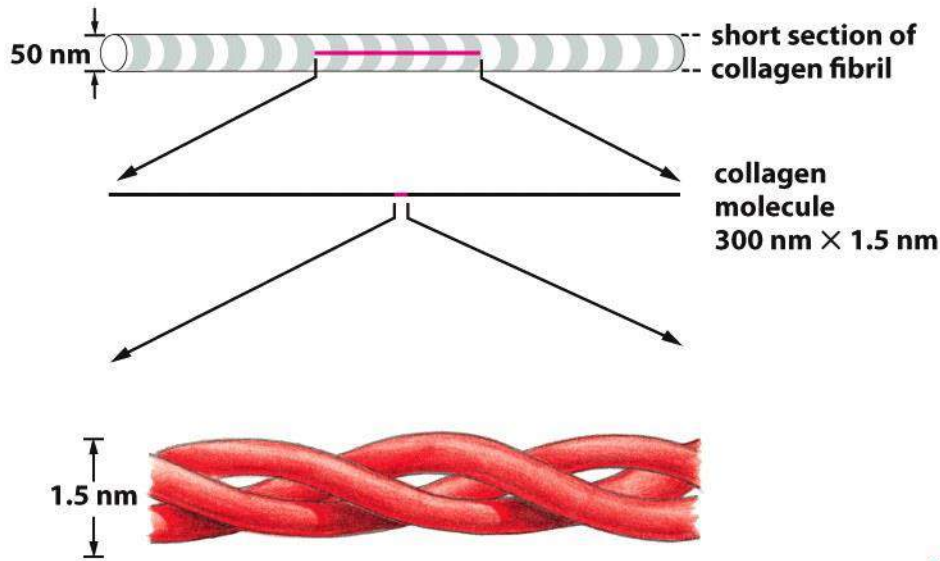
Figure 4-24 *Essential Cell Biology, 3rd ed.*
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ウイルスのキャプシド

微小管
 (α-チューブリンとβ-チューブリン)

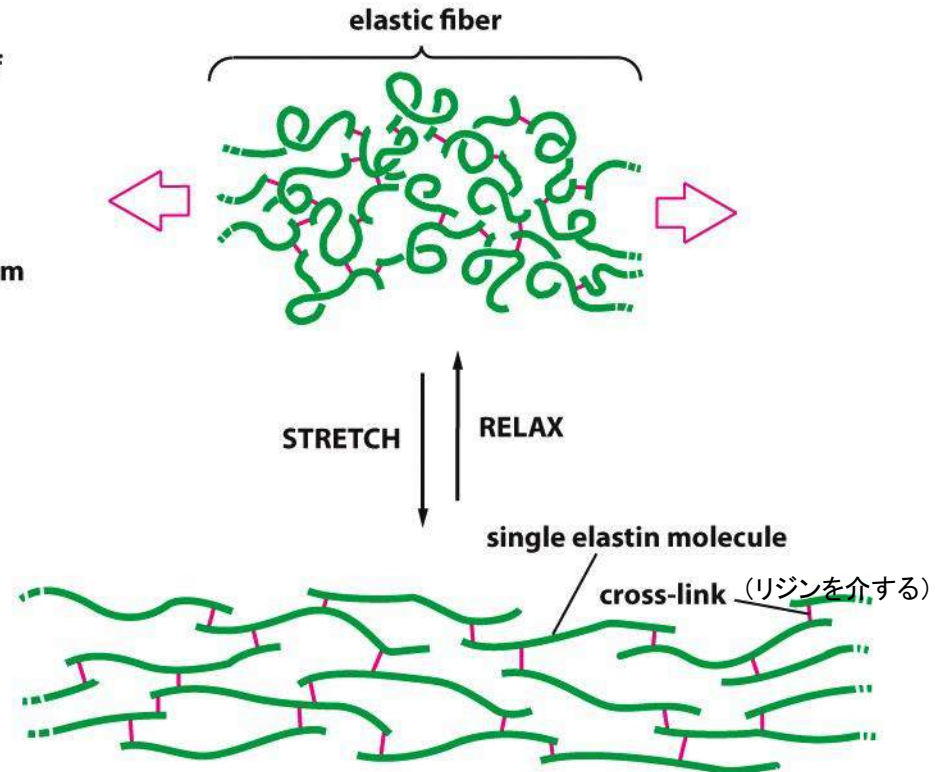
長い線維状のタンパク質

コラーゲン



(A)

エラスチン



(B)

皮膚、動脈、肺

タンパク質の架橋による安定化

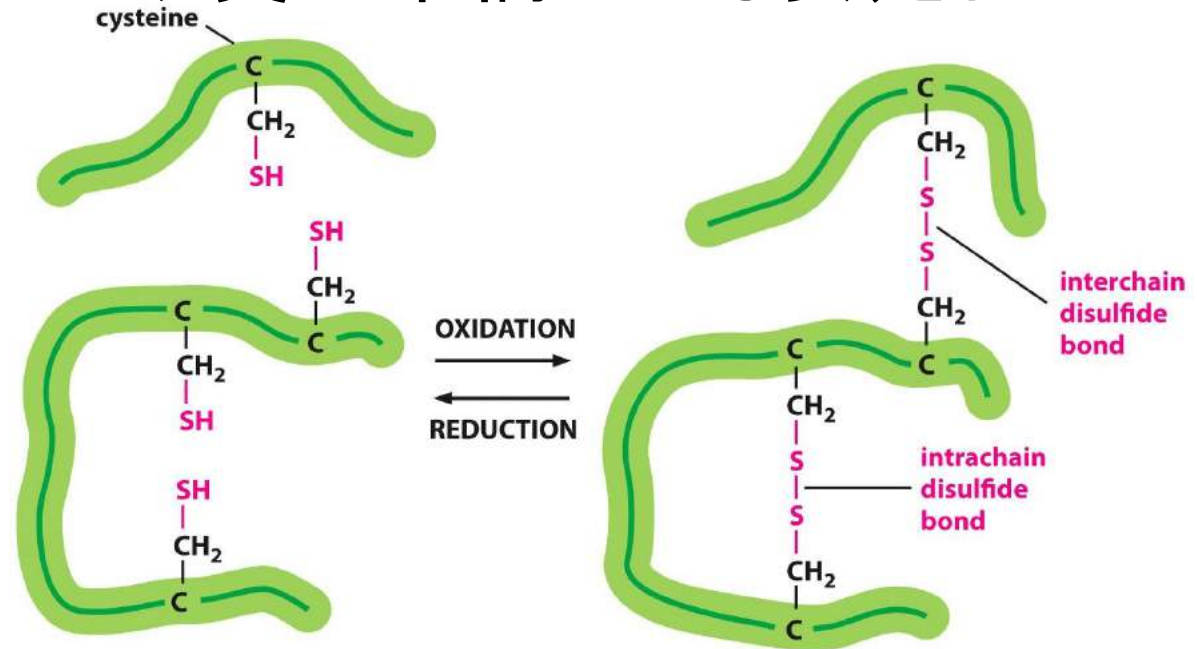
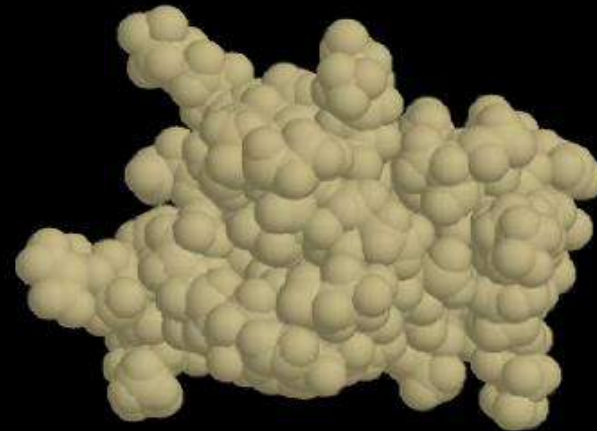


Figure 4-26 *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 139p.

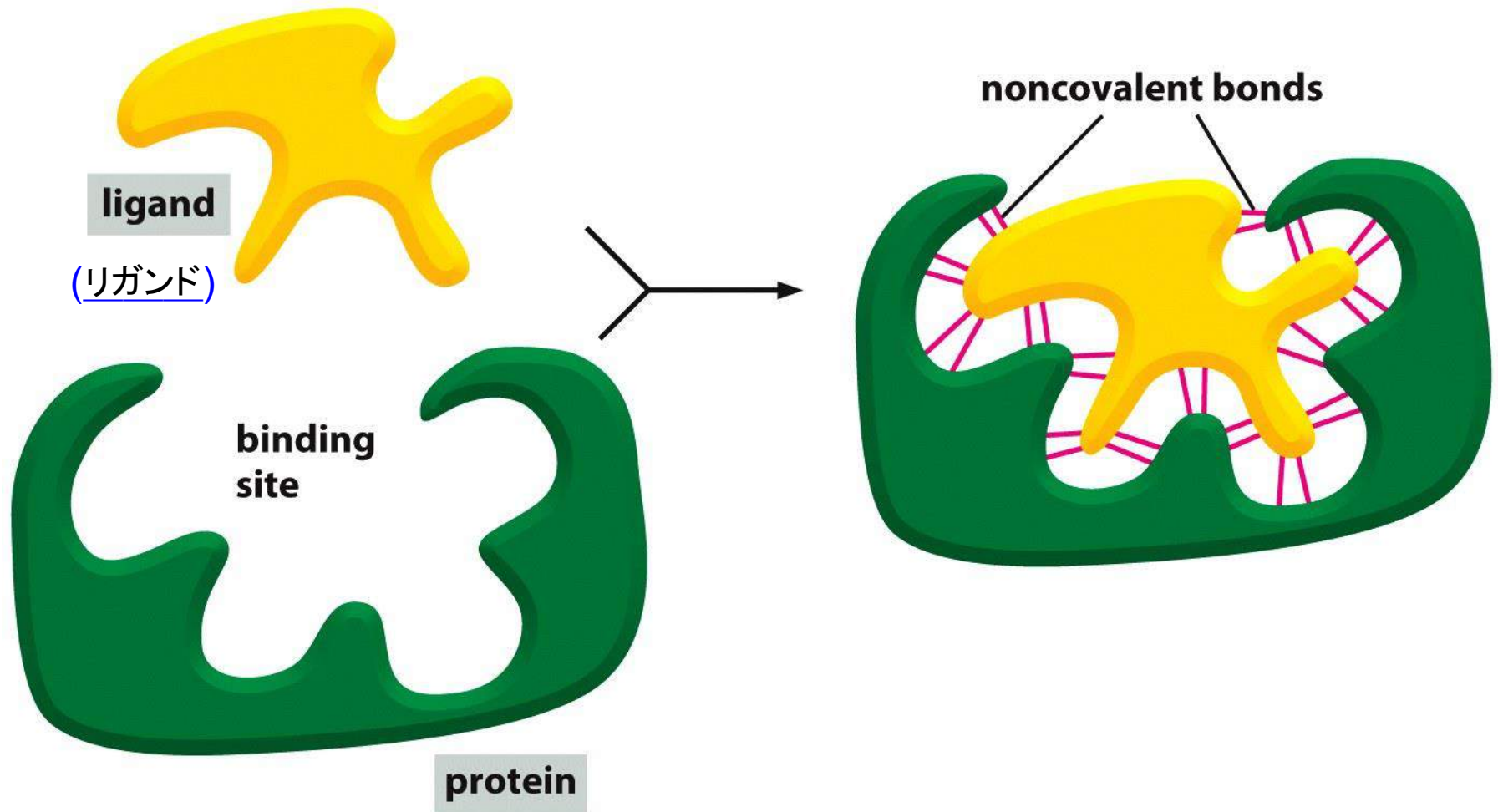
ジスルフィド結合

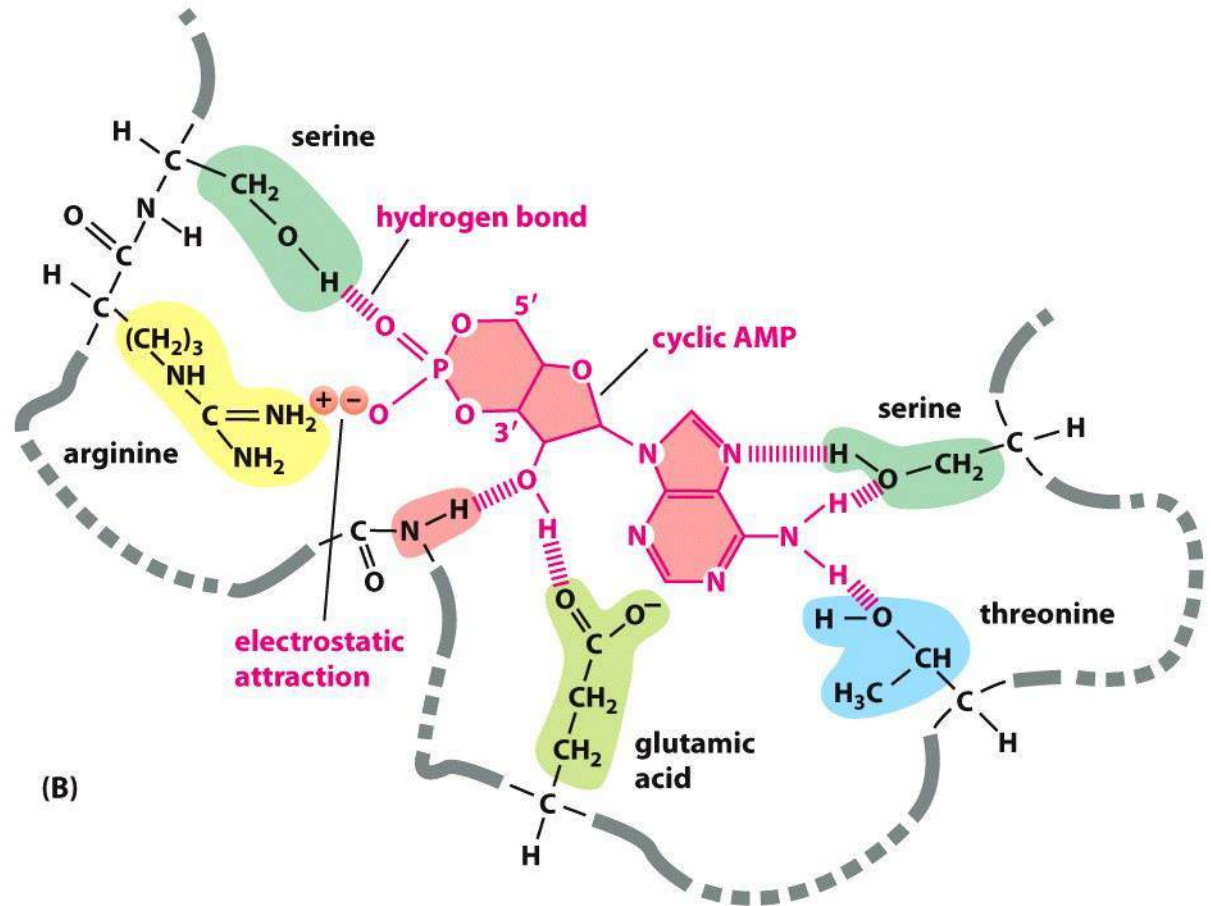
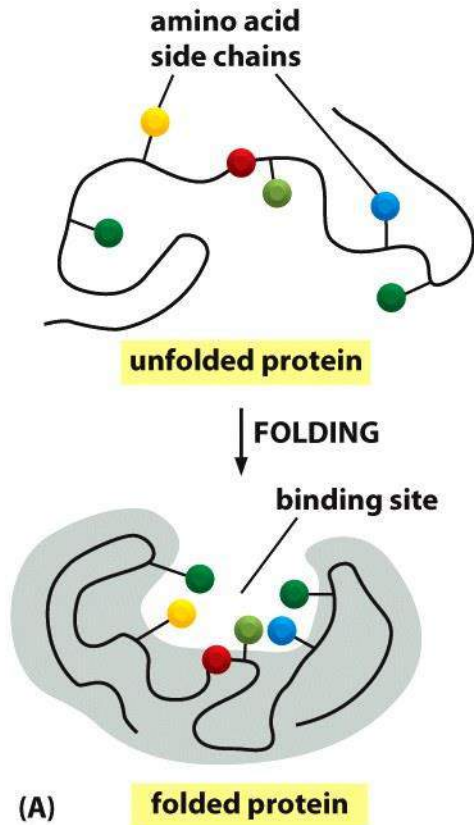
細胞外 に存在するタンパク質に特徴的



How proteins work

All proteins bind to other molecules





多数の 非共有結合 が相互作用に関与

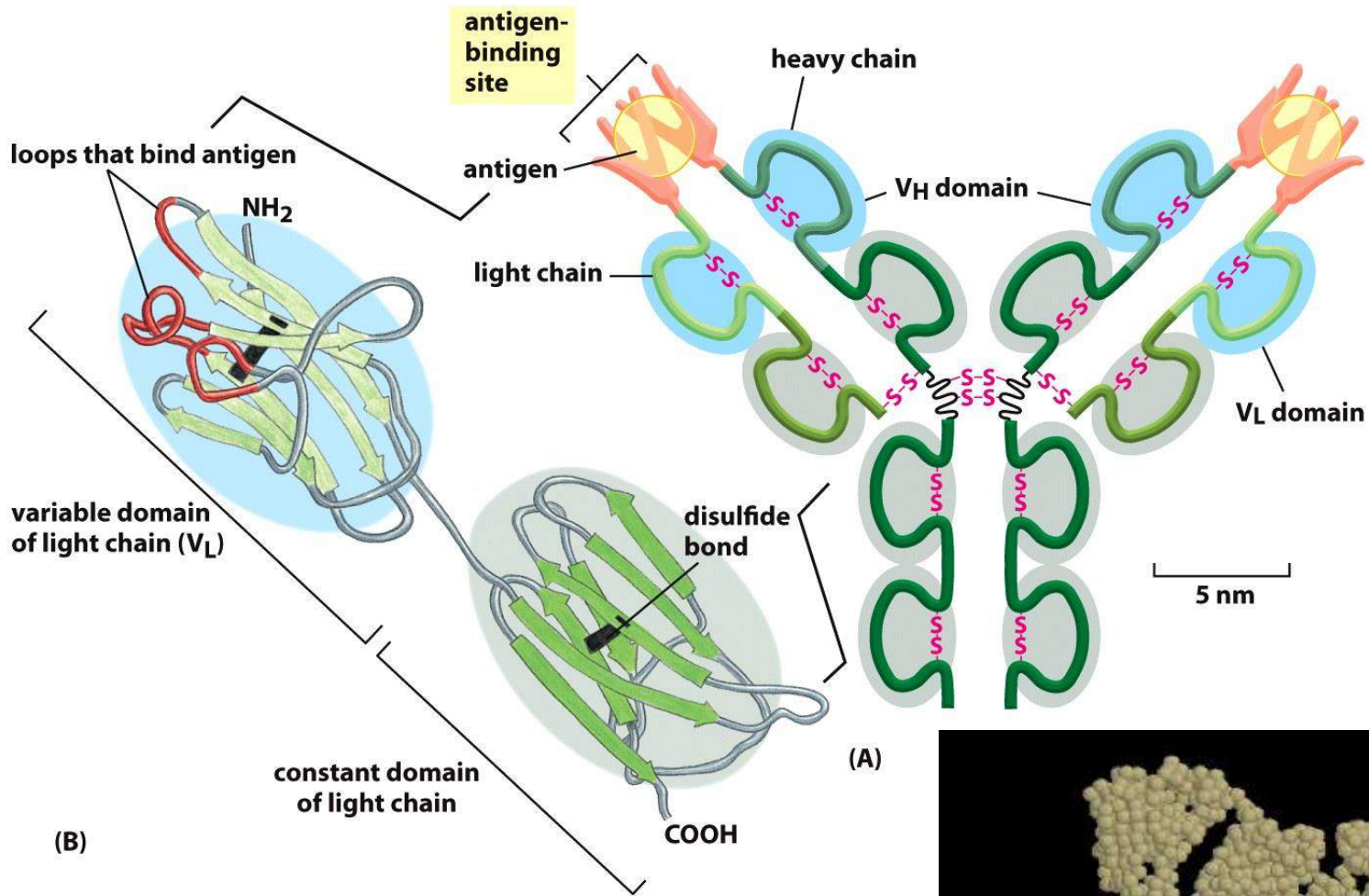
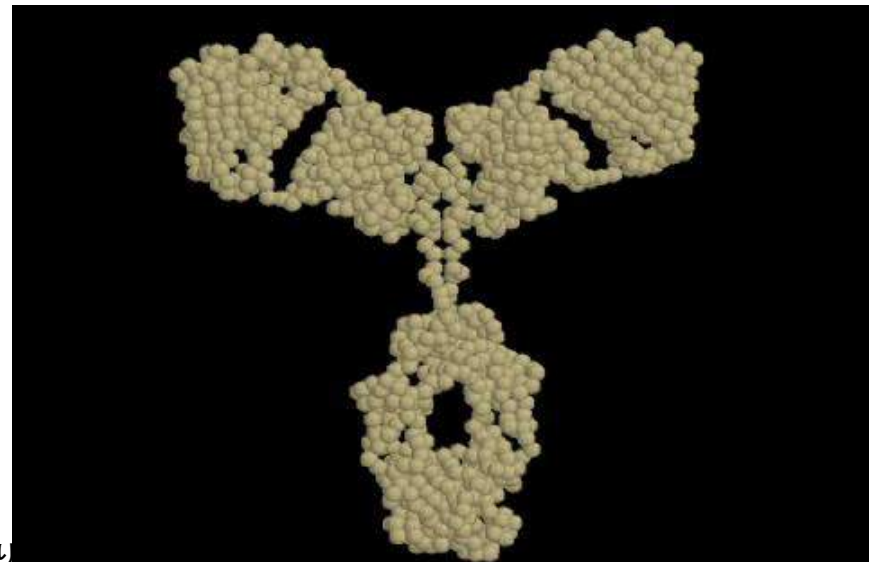


Figure 4-29 *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 142p.

Binding sites of antibodies are especially versatile (多目的)



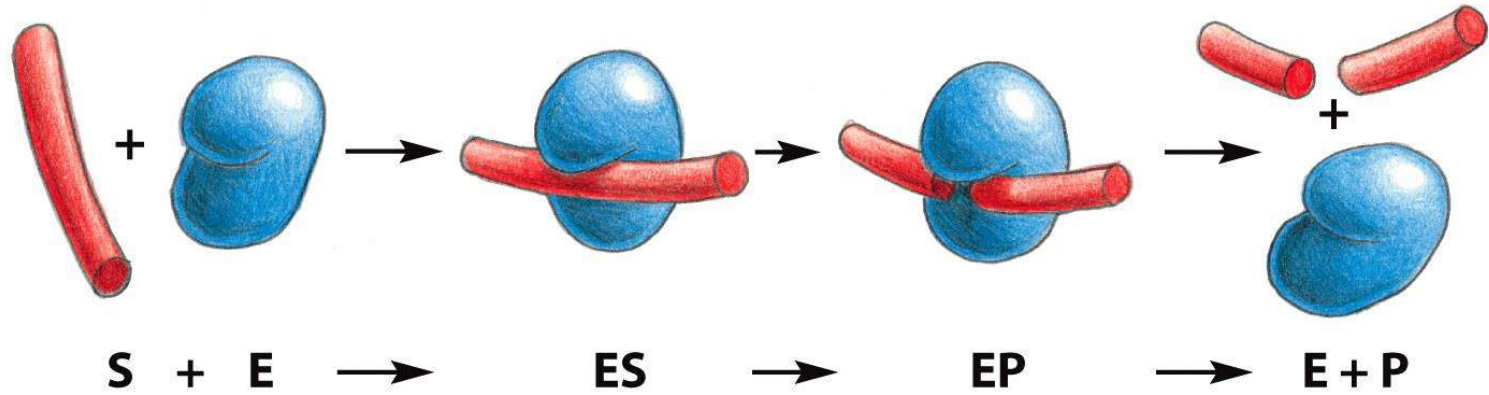
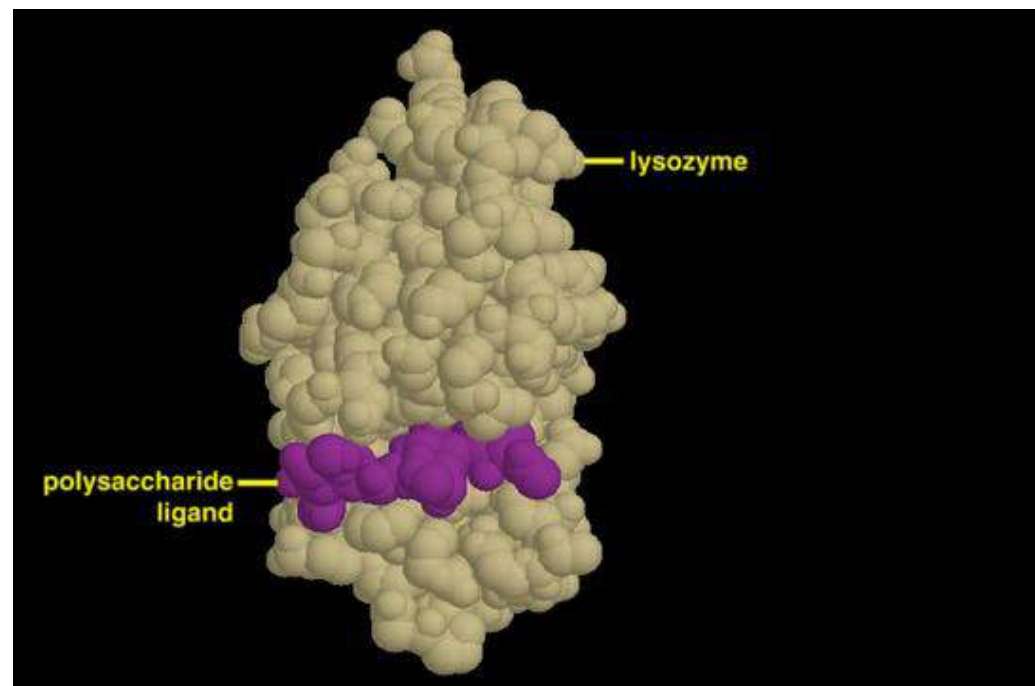
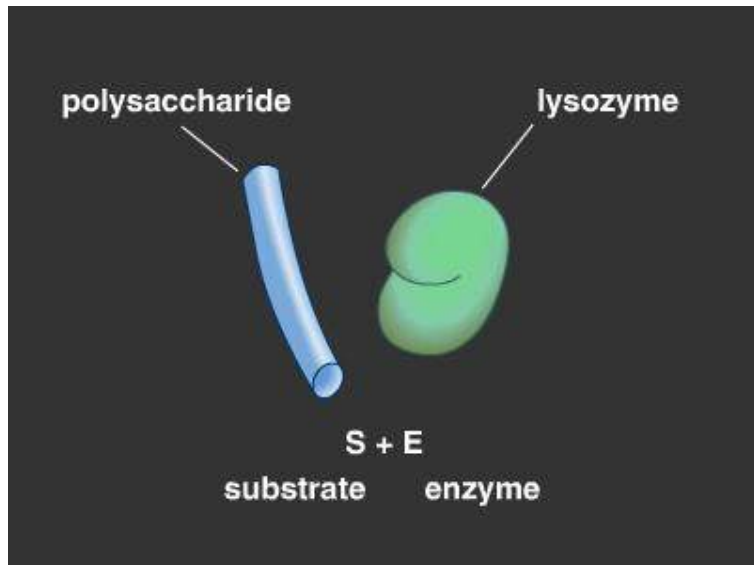
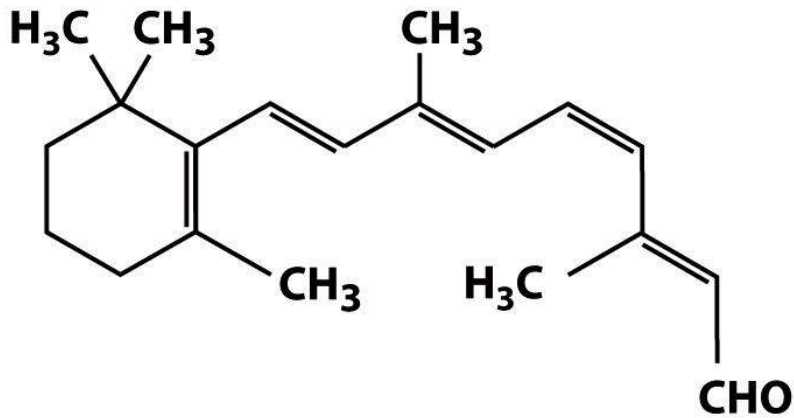


Figure 4-30(A) *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 146p.



タンパク質に強く結合している小分子が特別な機能を付加する

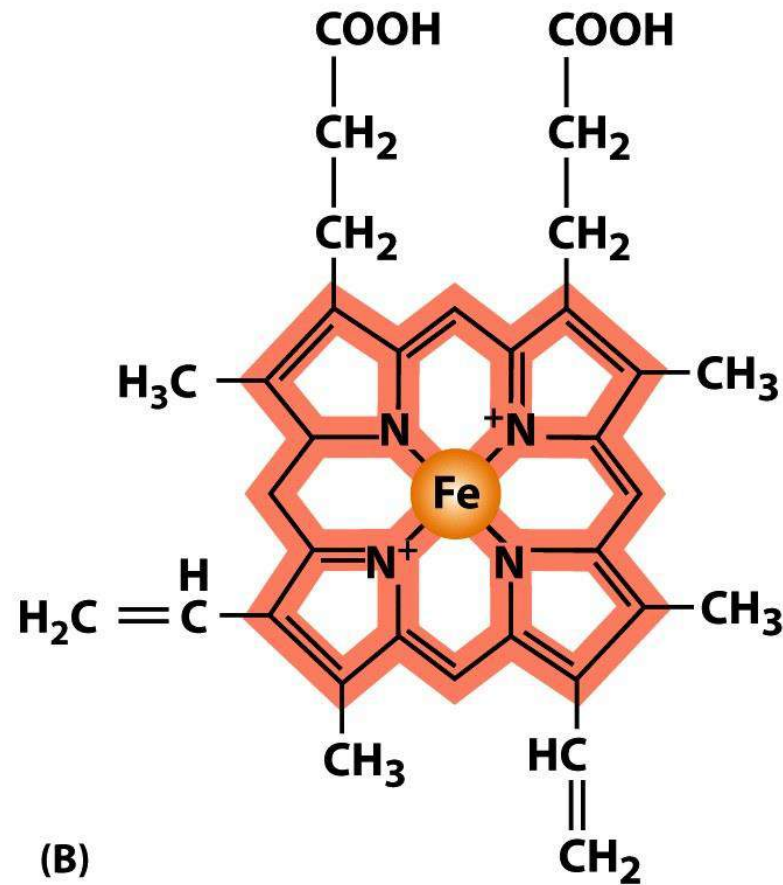


(A)

レチナール

ロドプシン

(7回膜貫通タンパク質)

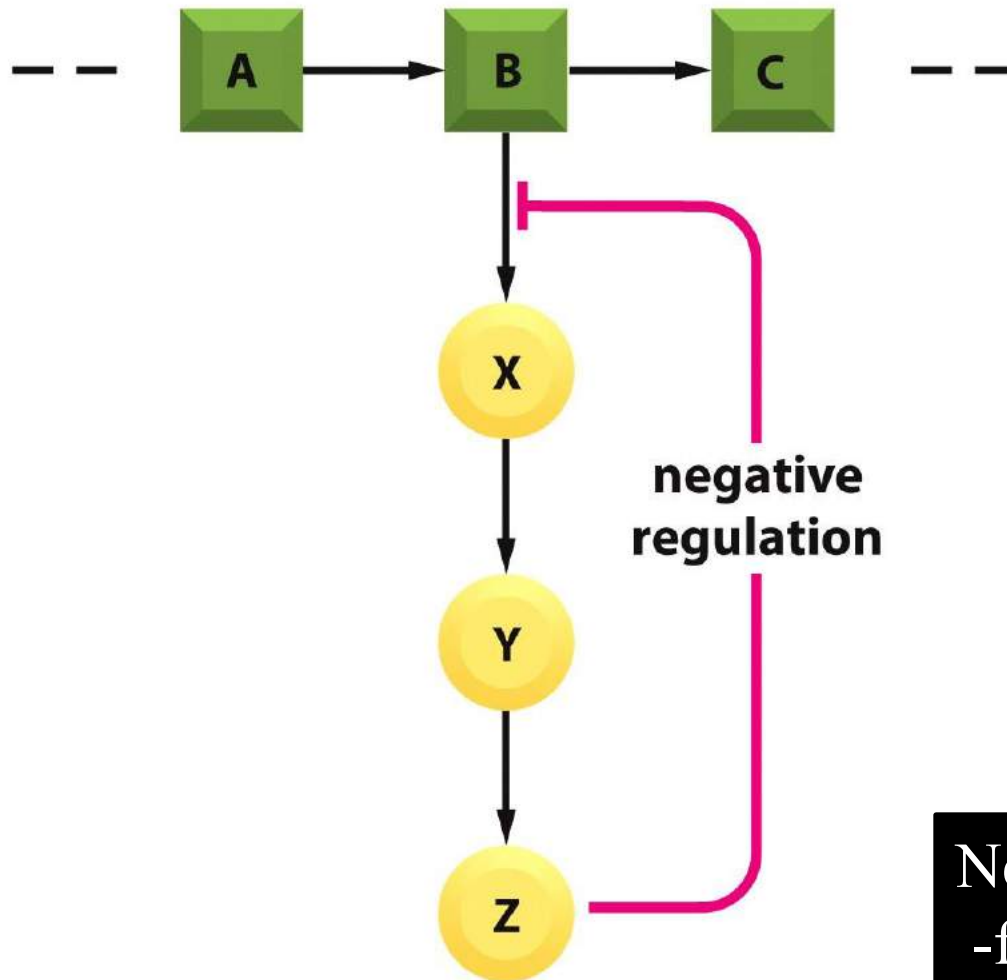


(B)

ヘム

ヘモグロビン

How proteins are controlled

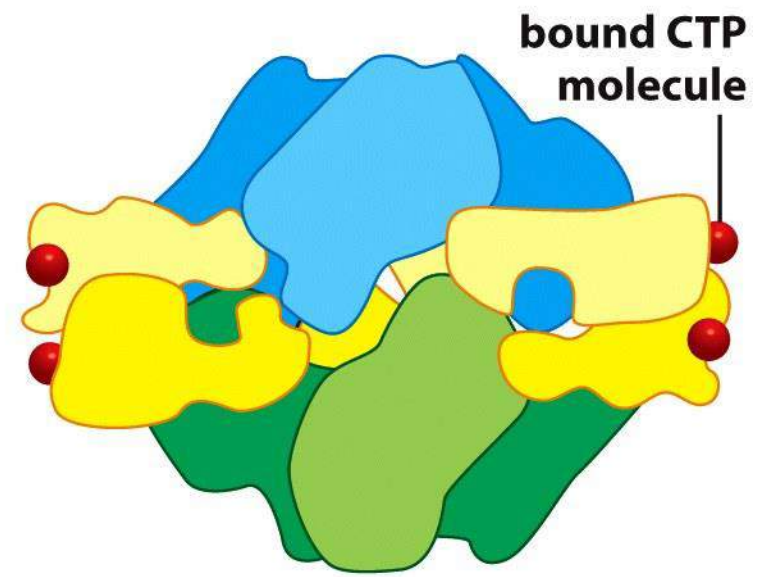
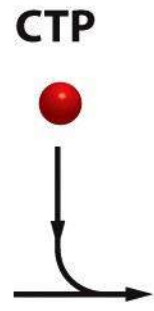
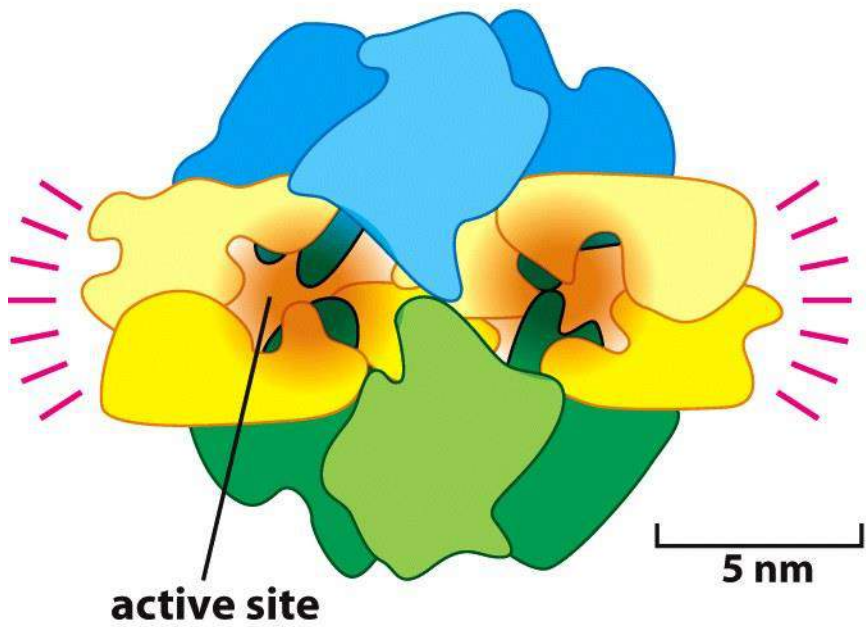


Feedback inhibition regulates the flow through biosynthetic pathways

Negative regulation
-feedback inhibition
Positive regulation

ON

OFF



ACTIVE ENZYME

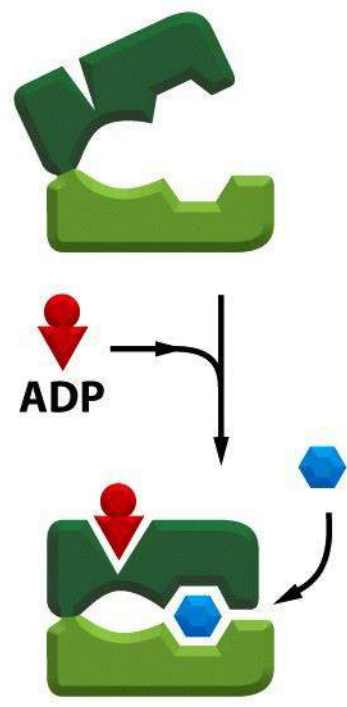
INACTIVE ENZYME

大腸菌のアスパラギン酸カルバモイル基転移酵素
 シトシン3リン酸(CTP)が最終生成物

Figure 4-36 *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 152p.

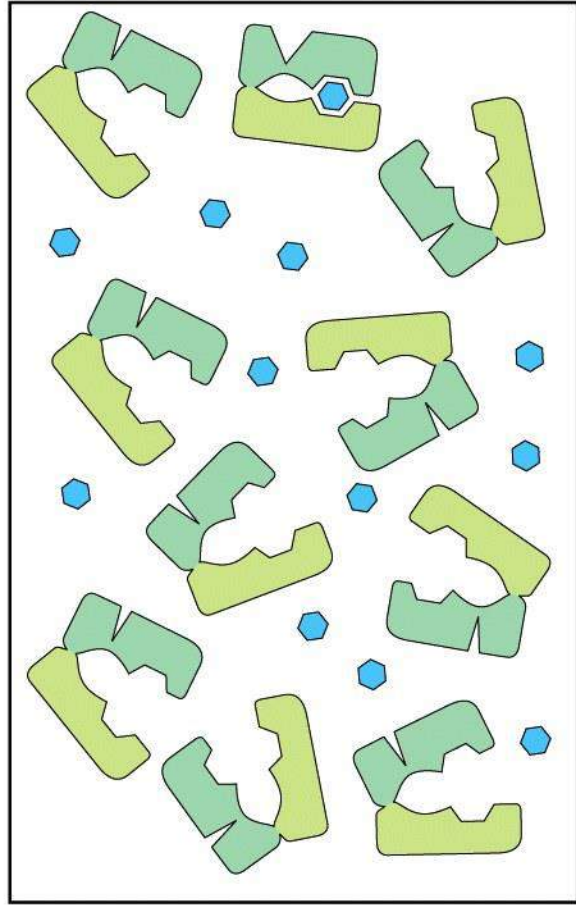
Allosteric Enzyme
 -regulatory molecule
 -substrate

INACTIVE



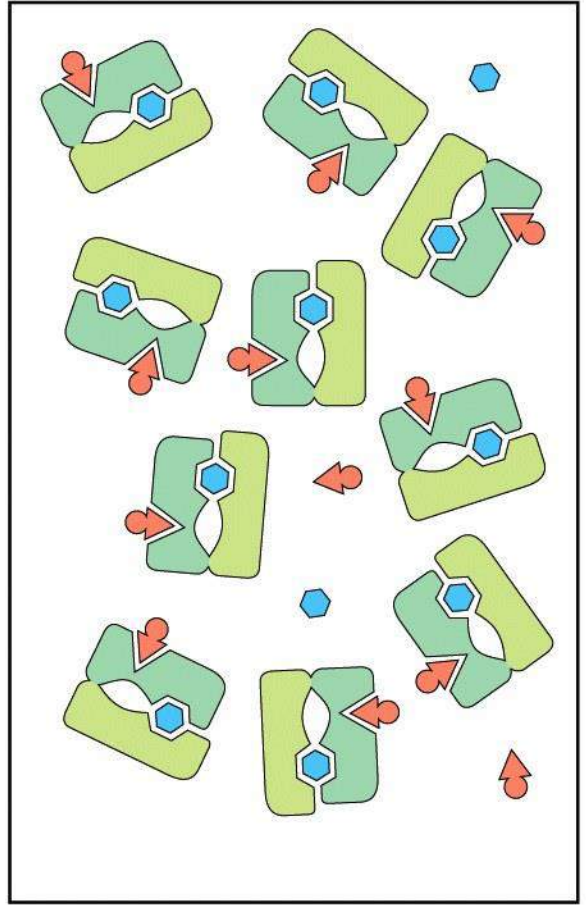
ACTIVE

(A)



(B) 10% active

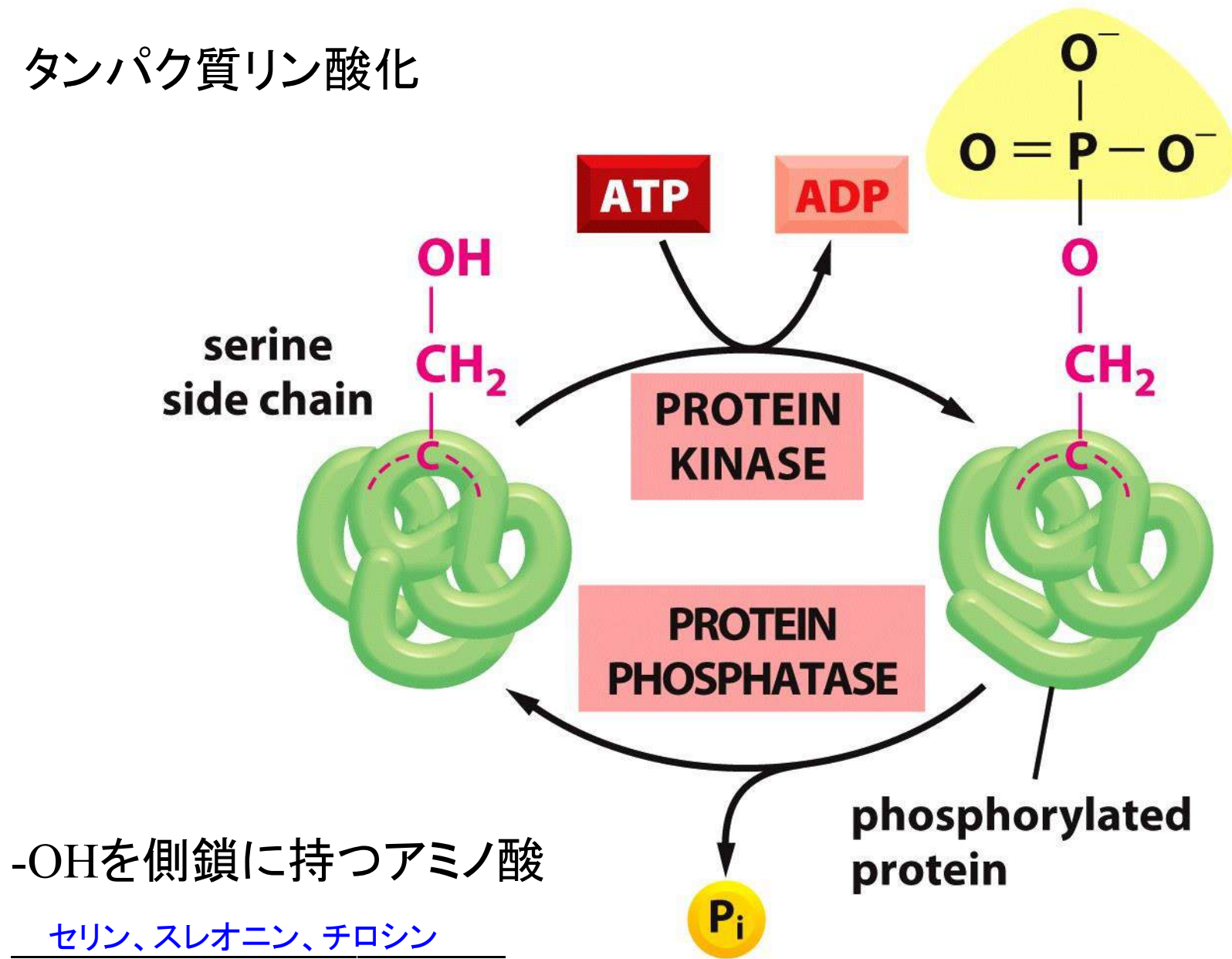
ADP
positive feedback



(C) 100% active

糖の異化に関わる仮想の酵素

タンパク質リン酸化



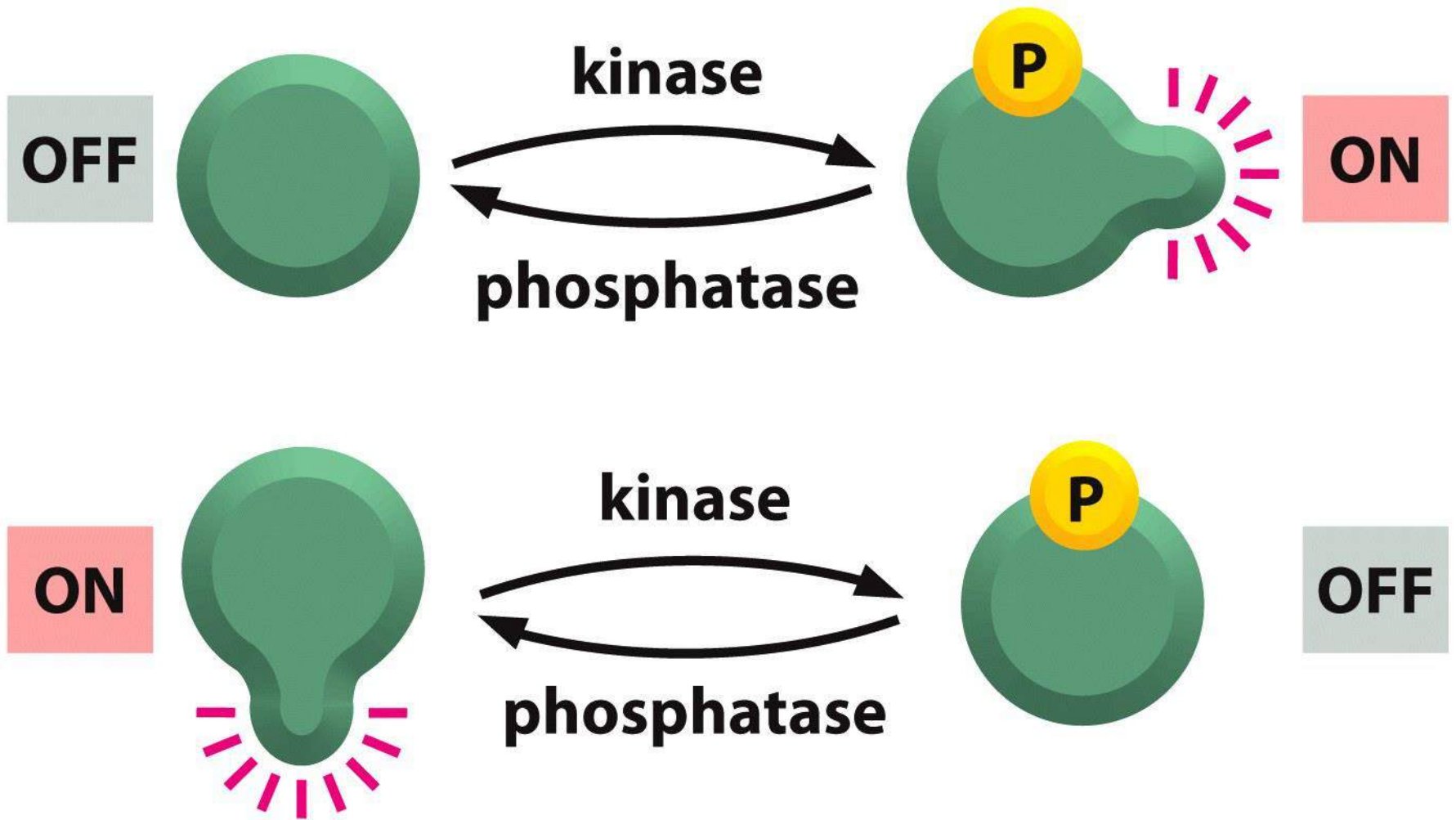


Figure 4-38(B) *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 153p.

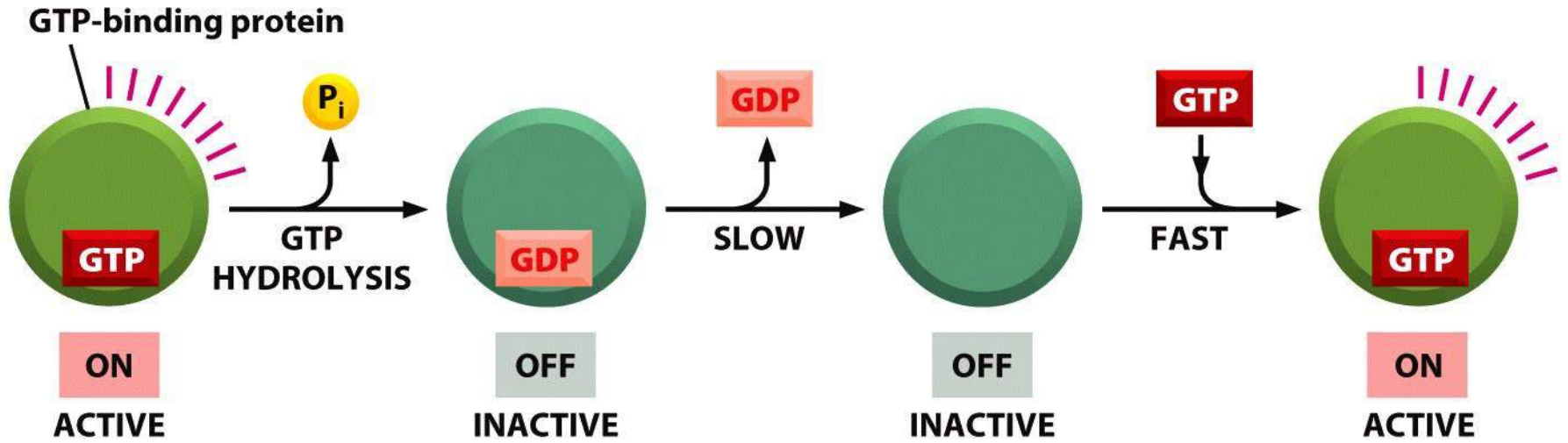
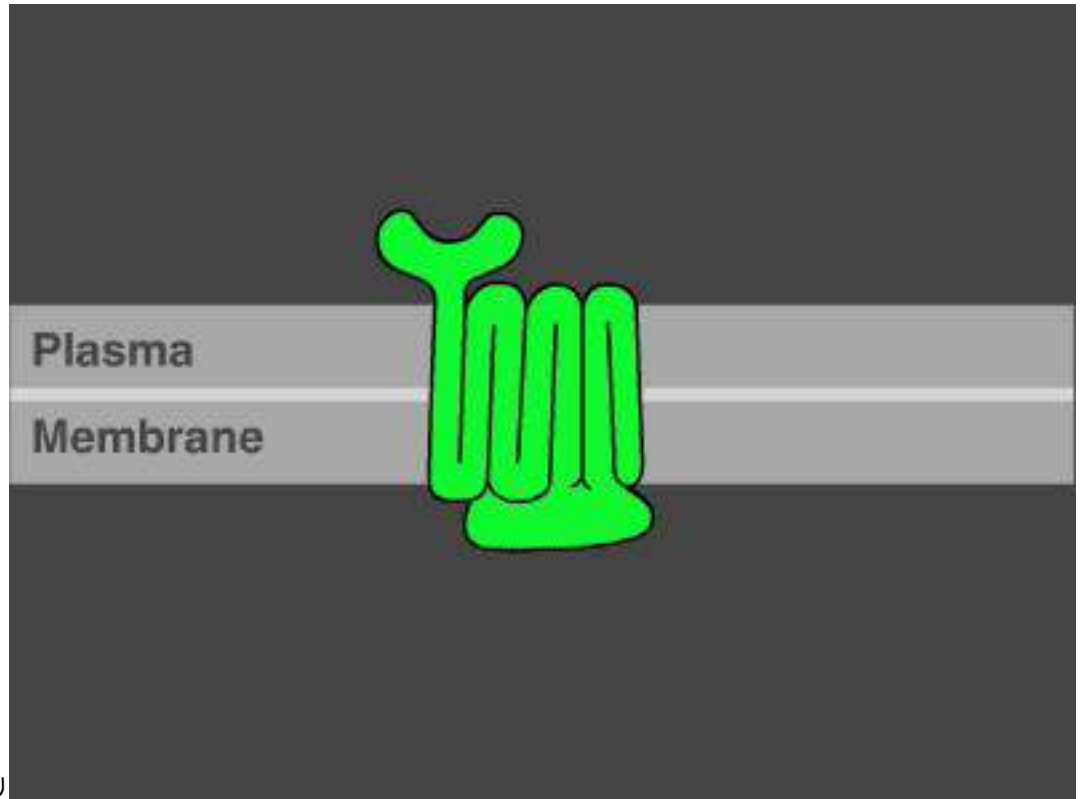


Figure 4-39 *Essential Cell Biology*, 3rd ed. (© Garland Science 2010) 153p.



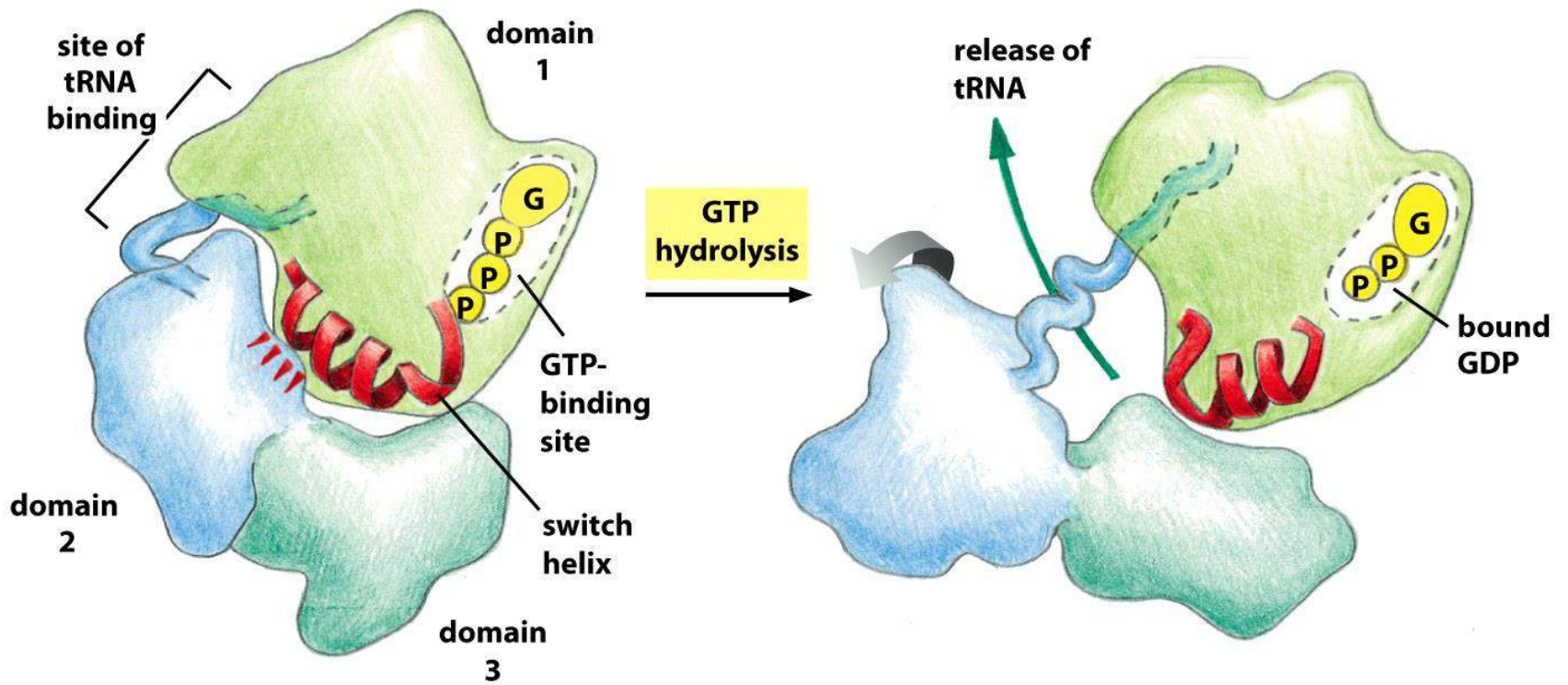
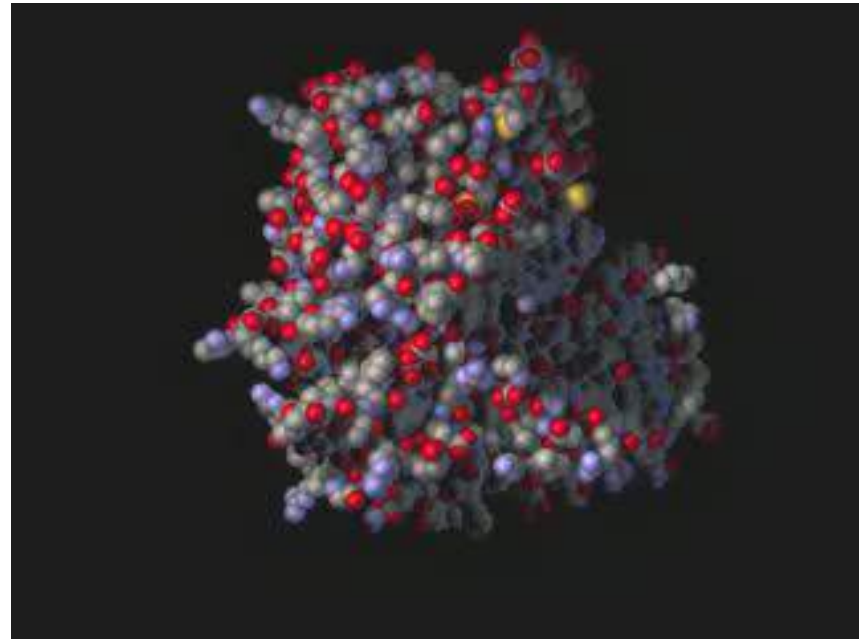


Figure 4-40 *Essential Cell Biology, 3rd ed.* (© Garland Science 2010) 154p.

細菌の伸長因子Tu (EF-Tu)

真核生物のEF1に相当

アミノアシルtRNA、GTPと結合し、リボソームのA部位にアミノアシルtRNAを供給する。



分子構造解析手法

X線結晶構造解析法

X線溶液散乱法 (SAXS)

核磁気共鳴法 (NMR)

分子動力学シミュレーション