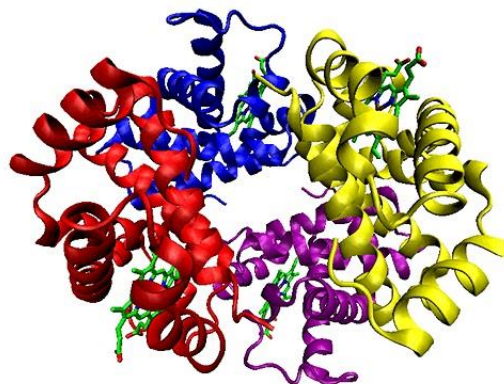


Lecture 7-2

Biomolecules - Protein



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Recommended Textbook:

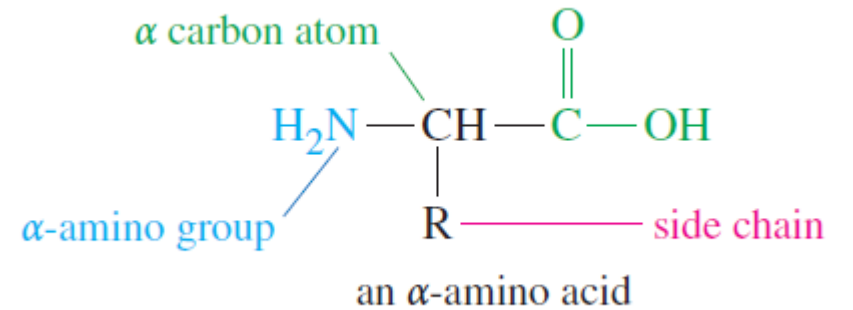
Chapter 23 and 25 in *Organic Chemistry*, 8th Edition, L. G. Wade, Jr., **2010**, Prentice Hall (Pearson Education)

Protein

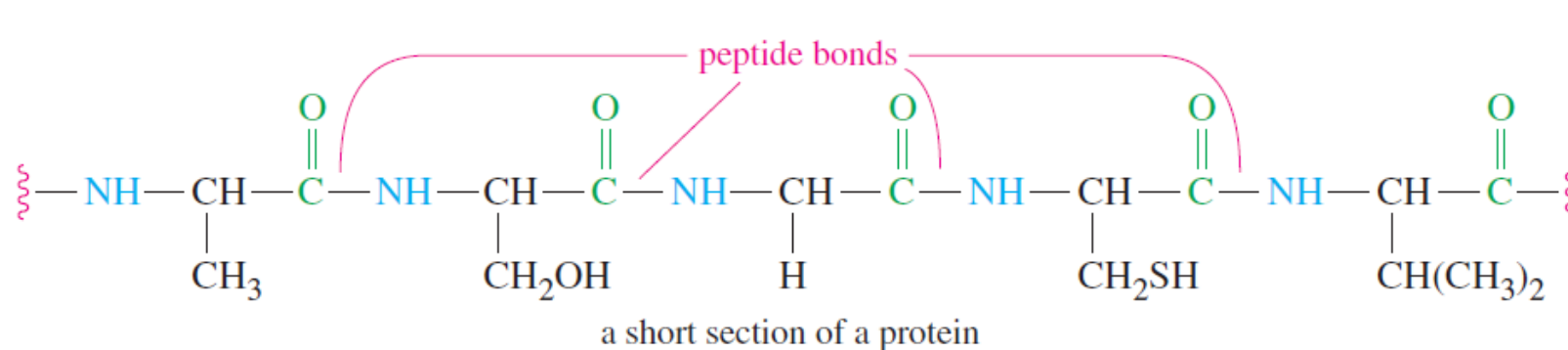
- Proteins are the **most abundant** organic molecules in **animals**, playing important roles in all aspects of cell structure and function



- Proteins are **biopolymers** of **α -amino acids**, so named because the amino group is bonded to the α carbon atom, next to the carbonyl group

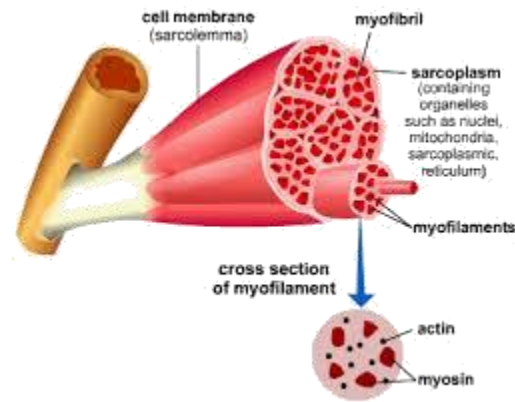
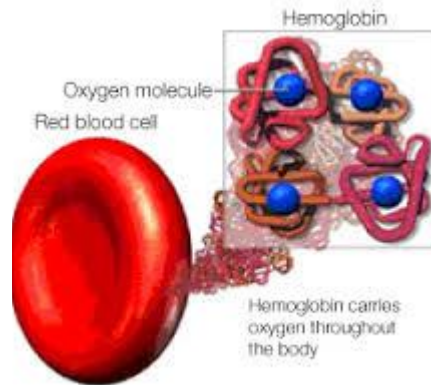
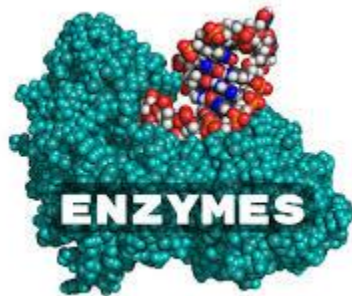


- The individual amino acid subunits are joined by **amide linkages** called **peptide bonds**



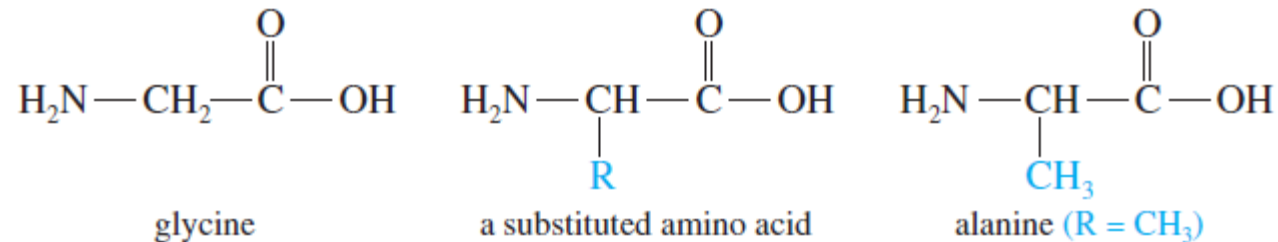
Protein – Examples

Class of Protein	Example	Function of Example
structural proteins	collagen, keratin	strengthen tendons, skin, hair, nails
enzymes	DNA polymerase	replicates and repairs DNA
transport proteins	hemoglobin	transports O ₂ to the cells
contractile proteins	actin, myosin	cause contraction of muscles
protective proteins	antibodies	complex with foreign proteins
hormones	insulin	regulates glucose metabolism
toxins	snake venoms	incapacitate prey



- The term amino acid might mean any molecule containing both an **amino group** and any type of **acid group**; however, the term is almost always used to refer to an **α -amino carboxylic acid**

- The simplest α -amino acid is **aminoacetic acid**, called **glycine**. Other common amino acids have **side chains** (symbolised by **R**) substituted on the **α carbon**; for example, **alanine** is the amino acid with a **methyl side chain**



Standard α -Amino Acids

20 standard amino acids, grouped according to the chemical properties of side chains

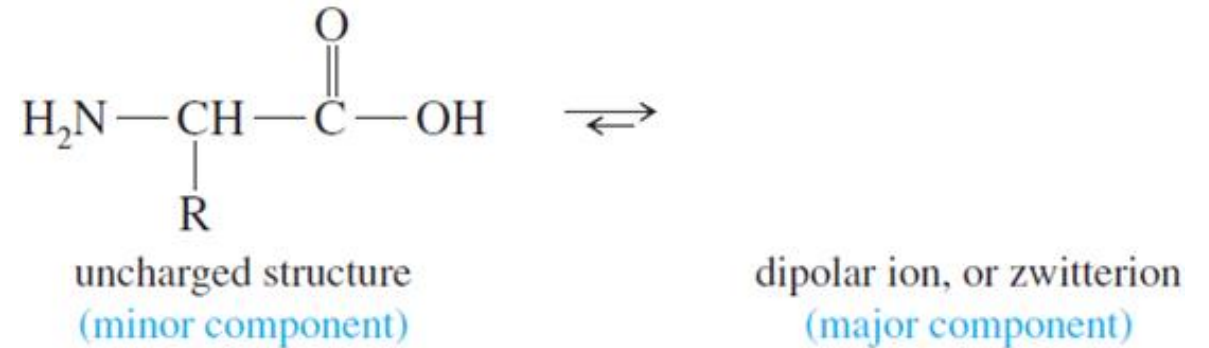
Name	Symbol	Abbreviation	Structure	Functional Group in Side Chain	Isoelectric Point
<i>side chain is nonpolar, H or alkyl</i>					
glycine	G	Gly	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{H} \end{array}$	none	6.0
alanine	A	Ala	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_3 \end{array}$	alkyl group	6.0
*valine	V	Val	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH} \\ / \quad \backslash \\ \text{CH}_3 \quad \text{CH}_3 \end{array}$	alkyl group	6.0
*leucine	L	Leu	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{CH}-\text{CH}_3 \\ \\ \text{CH}_3 \end{array}$	alkyl group	6.0
*isoleucine	I	Ile	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_3-\text{CH}-\text{CH}_2\text{CH}_3 \end{array}$	alkyl group	6.0
*phenylalanine	F	Phe	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{C}_6\text{H}_5 \end{array}$	aromatic group	5.5
proline	P	Pro	$\begin{array}{c} \text{HN}-\text{CH}-\text{COOH} \\ / \quad \backslash \\ \text{H}_2\text{C} \quad \text{CH}_2 \\ \\ \text{CH}_2 \end{array}$	rigid cyclic structure	6.3
<i>*essential amino acid</i>					

Name	Symbol	Abbreviation	Structure	Functional Group in Side Chain	Isoelectric Point
side chain contains an —OH					
serine	S	Ser	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{OH} \end{array}$	hydroxyl group	5.7
*threonine	T	Thr	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{HO}-\text{CH}-\text{CH}_3 \end{array}$	hydroxyl group	5.6
tyrosine	Y	Tyr	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{C}_6\text{H}_4-\text{OH} \end{array}$	phenolic—OH group	5.7
side chain contains sulfur					
cysteine	C	Cys	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{SH} \end{array}$	thiol	5.0
*methionine	M	Met	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{CH}_2-\text{S}-\text{CH}_3 \end{array}$	sulfide	5.7
side chain is acidic					
aspartic acid	D	Asp	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{COOH} \end{array}$	carboxylic acid	2.8
glutamic acid	E	Glu	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{CH}_2-\text{COOH} \end{array}$	carboxylic acid	3.2

Name	Symbol	Abbreviation	Structure	Functional Group in Side Chain	Isoelectric Point
side chain is basic					
*lysine	K	Lys	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{CH}_2-\text{CH}_2-\text{CH}_2-\text{NH}_2 \end{array}$	amino group	9.7
*arginine	R	Arg	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{CH}_2-\text{CH}_2-\text{NH}-\text{C}-\text{NH}_2 \\ \\ \text{NH} \end{array}$	guanidino group	10.8
*histidine	H	His	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2 \\ \\ \text{Imidazole ring} \end{array}$	imidazole ring	7.6
side chain contains nonbasic nitrogen					
asparagine	N	Asn	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{C}-\text{NH}_2 \\ \\ \text{O} \end{array}$	amide	5.4
glutamine	Q	Gln	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2-\text{CH}_2-\text{C}-\text{NH}_2 \\ \\ \text{O} \end{array}$	amide	5.7
*tryptophan	W	Trp	$\begin{array}{c} \text{H}_2\text{N}-\text{CH}-\text{COOH} \\ \\ \text{CH}_2 \\ \\ \text{Indole ring} \end{array}$	indole	5.9

α -Amino Acids – Acid-Base Properties

- Although we commonly write amino acids with an intact carboxyl $-\text{COOH}$ group and amino $-\text{NH}_2$ group, their actual structure is **ionic** and **depends on the pH**
- The carboxyl group loses a proton, giving a **carboxylate ion**, and the amino group is protonated to an **ammonium ion**. This structure is called a **zwitterion**, giving them some **unusual properties**

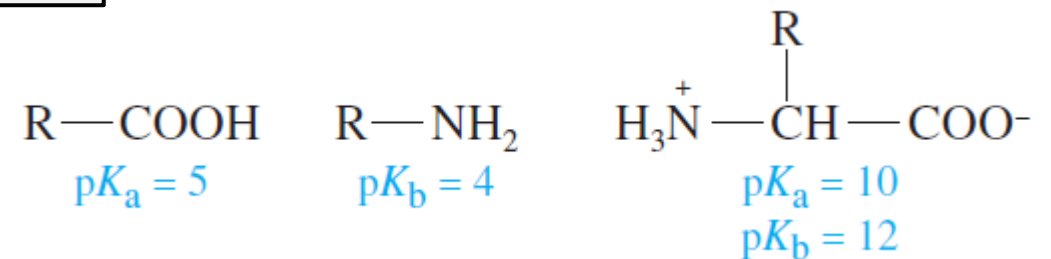


1) More **soluble in water** than they are in common organic solvents

2) **high melting points**, generally over $200\text{ }^\circ\text{C}$

3) much **larger dipole moments** than simple amines or simple acids

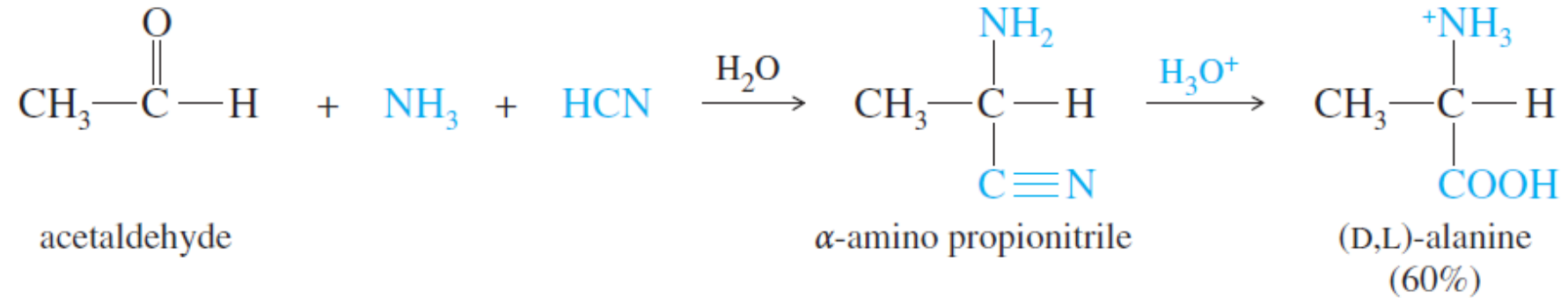
4) **Less acidic than most carboxylic acids** and **less basic than most amines**



α -Amino Acids – Synthesis

The Strecker Synthesis; formation of amino acids from appropriate aldehydes

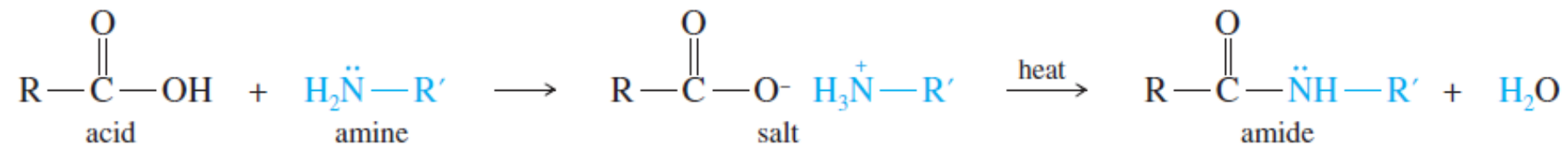
The Strecker synthesis of alanine



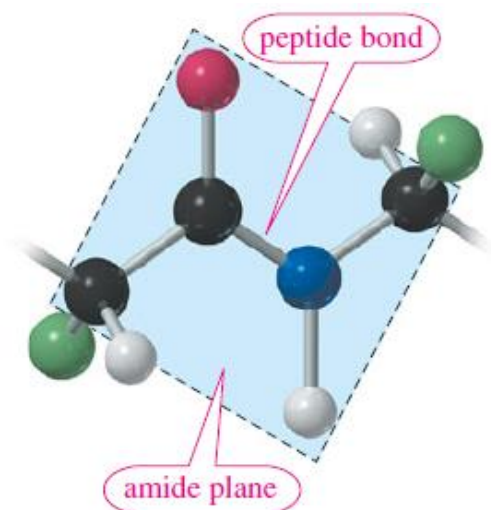
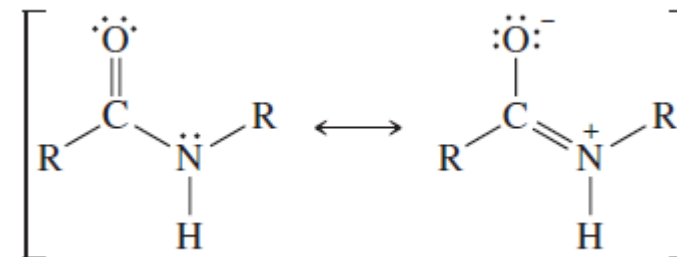
Mechanism:

Peptides

- Amines** and **acids** can condense, with the loss of water, to form **amides**



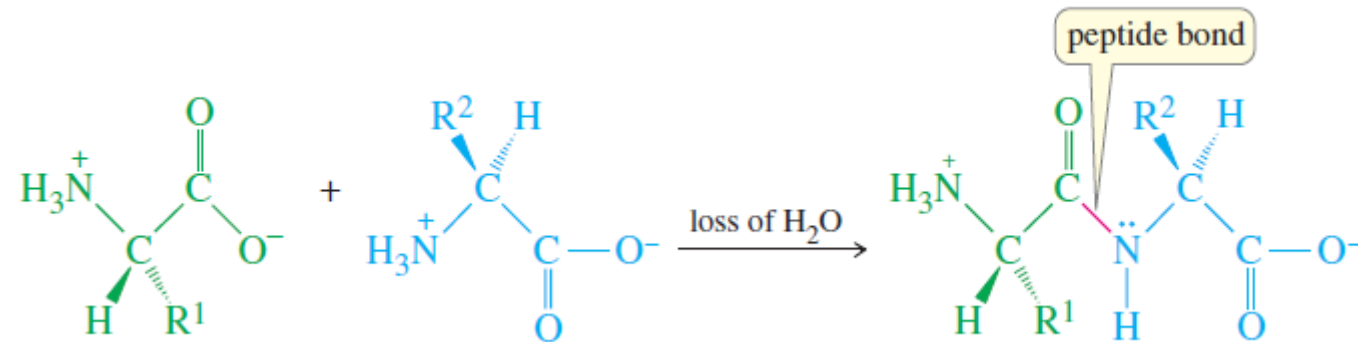
- Amides are the **most stable** acid derivatives. This stability is partly due to the strong **resonance interaction** between the nonbonding electrons on nitrogen and the carbonyl group



- The amide linkage between the amino acids is called a **peptide bond**

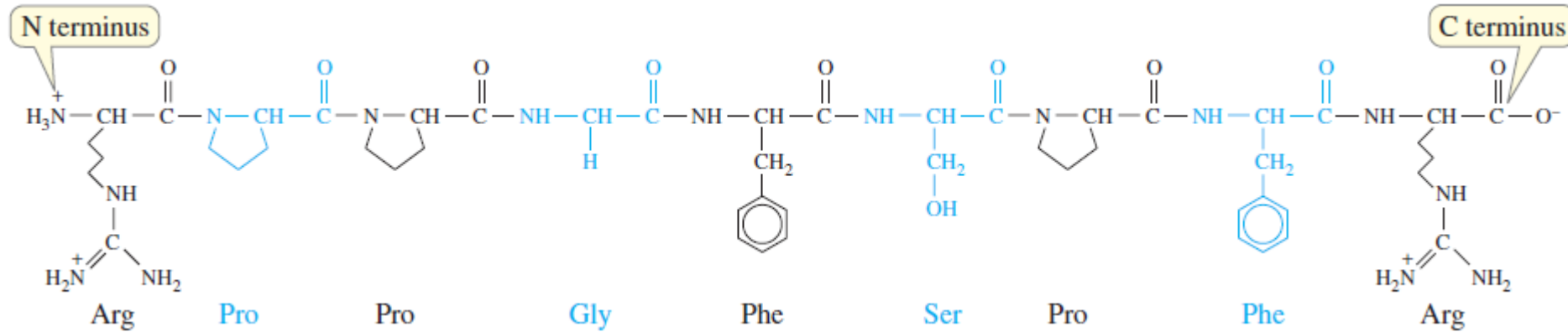
Peptides

- A **peptide** is a compound containing two or more amino acids linked by amide bonds
- Under the proper conditions, the amino group of one amino acid condenses with the carboxyl group of another
- The product is an amide called a **dipeptide** because it consists of two amino acids



- A **polypeptide** is a peptide containing many amino acid residues but usually having a molecular weight of less than about **5000**
- **Proteins** contain more amino acid units, with molecular weights ranging from about **5000** to about **40,000,000**

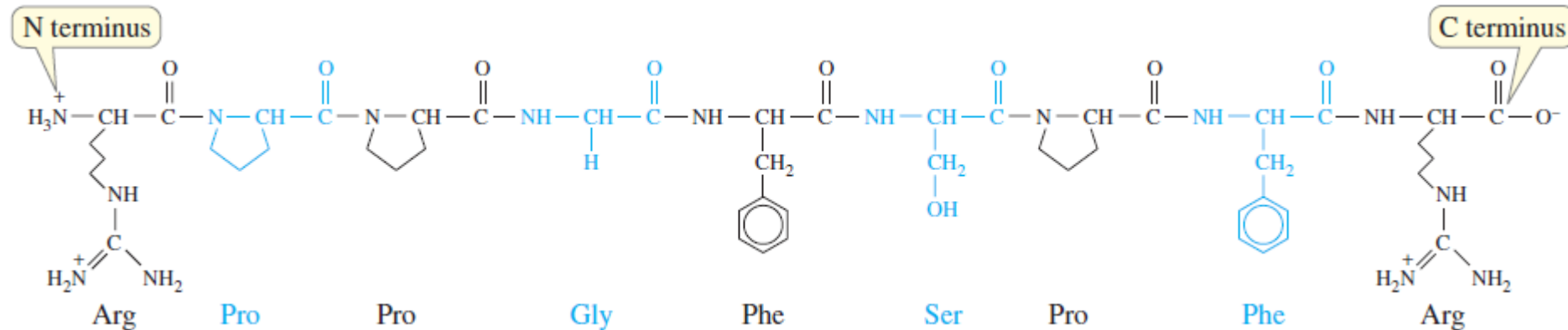
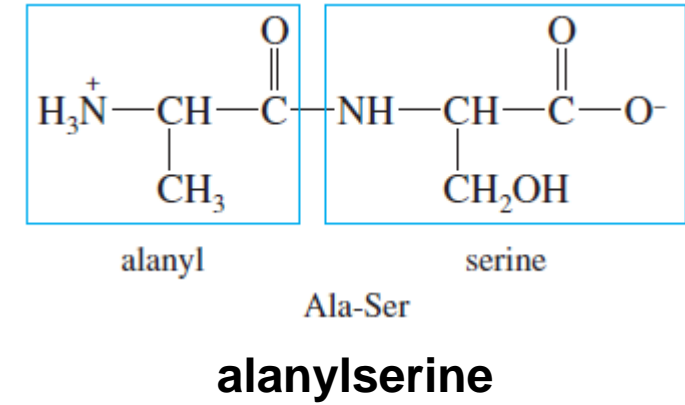
- Nonapeptide **bradykinin**, a human hormone that helps to control blood pressure



- The end of the peptide with the free amino group ($-\text{NH}_3^+$) is called the **N terminus**
- The end with the free carboxyl group ($-\text{CO}_2^-$) is called the **C terminus**
- Peptide structures are generally drawn with the **N terminus at the left** and the **C terminus at the right**

Peptides – Nomenclature

- The names of peptides reflect the names of the **amino acid residues** involved in the amide linkages, **beginning at the N terminus**; All **except the last** are given the **-yl suffix** of acyl groups



arginyl prolyl prolyl glycyll phenylalanyl seryl prolyl phenylalanyl arginine

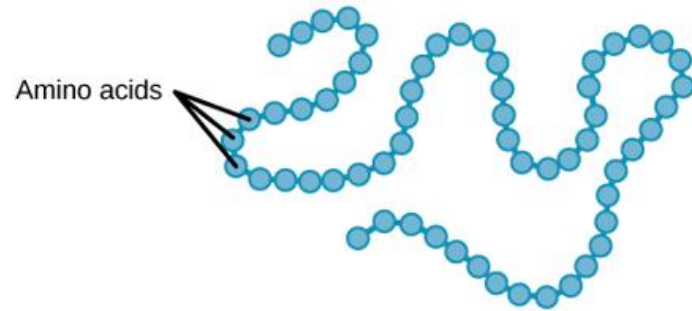
- A **shorthand system** is more convenient, representing each amino acid by its three-letter abbreviation : **Arg-Pro-Pro-Gly-Phe-Ser-Pro-Phe-Arg**
- Single-letter symbols** are becoming widely used as well : **RPPGFSPFR**

Proteins – Structure

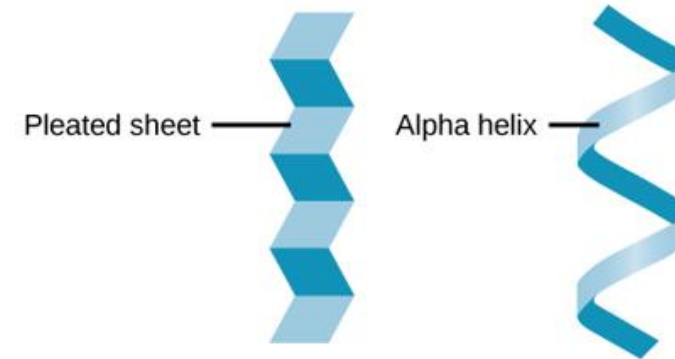
- **Proteins** contain more amino acid units, with molecular weights ranging from about **5000** to about **40,000,000**

Level of Protein Structure:

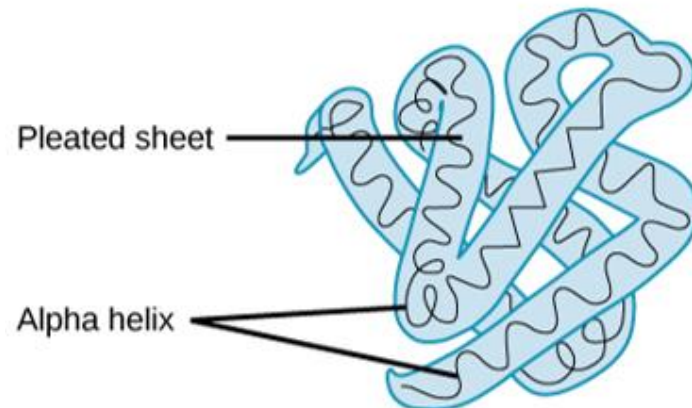
1) **Primary:** sequence of a chain of amino acids



2) **Secondary:** folding of peptide into a repeating pattern due to H-bonding of peptide backbone



3) **Tertiary:** three dimensional folding pattern due to side chain interaction



4) **Quaternary:** protein that consists of more than one peptide chain



Proteins – Classification

- Proteins may be classified according to their **chemical composition**, their **shape**, or their **function**
- By **chemical composition** : *simple* and *conjugated* proteins
 - *Simple proteins*: those that hydrolyse to give **only amino acids**
 - *Conjugated proteins*: bonded to a nonprotein **prosthetic group** such as a sugar, a nucleic acid, a lipid, or some other group

TABLE 24-3 Classes of Conjugated Proteins

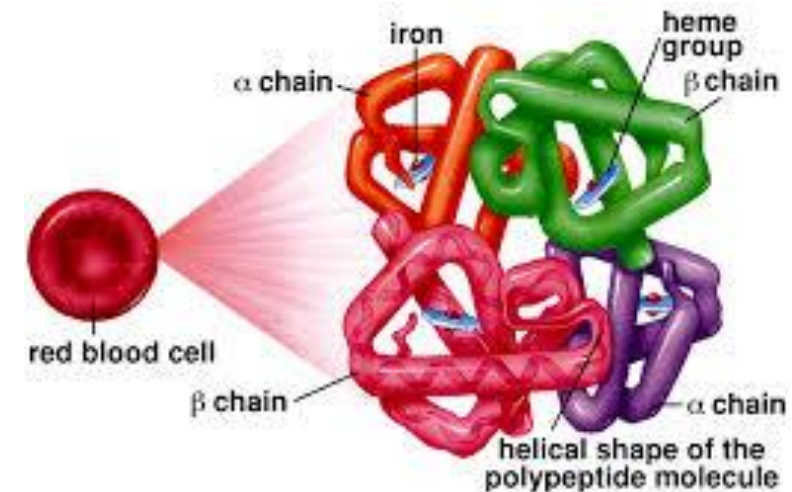
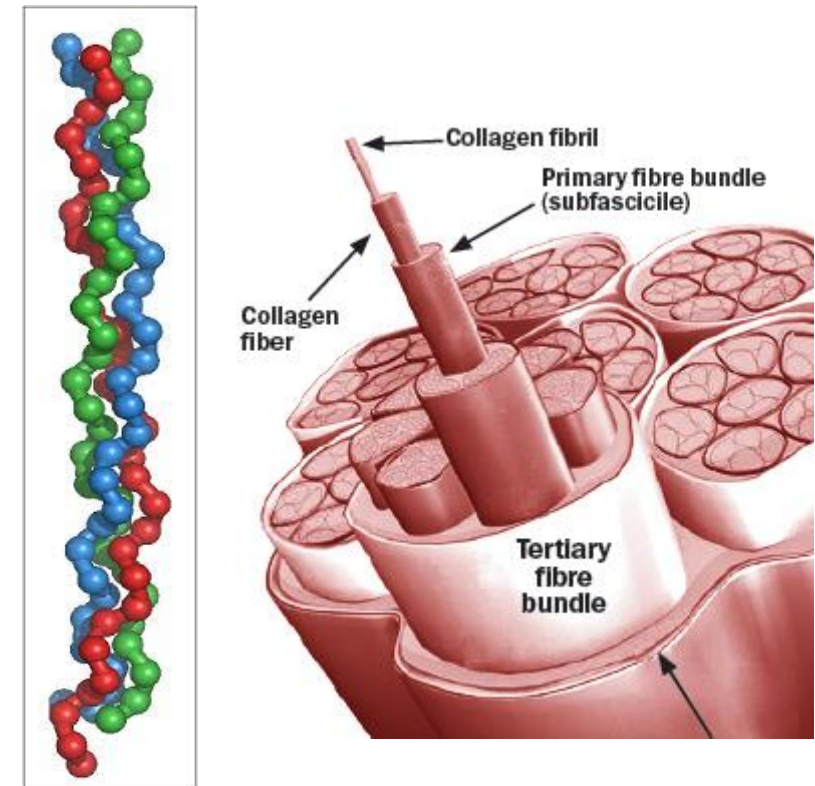
Class	Prosthetic Group	Examples
glycoproteins	carbohydrates	γ -globulin, interferon
nucleoproteins	nucleic acids	ribosomes, viruses
lipoproteins	fats, cholesterol	high-density lipoprotein
metalloproteins	a complexed metal	hemoglobin, cytochromes

Proteins – Classification

- By shape : *fibrous* and *globular* proteins
- **Fibrous proteins**: stringy, tough, and usually insoluble in water. They function primarily as **structural parts** of the organism

Examples : α -keratin in hooves and fingernails, and collagen in tendons
- **Globular proteins**: are folded into roughly spherical shapes. They usually function as **enzymes, hormones, or transport proteins**

Transport proteins bind to specific molecules and transport them in the blood or through the cell membrane; for example, **hemoglobin** transports oxygen in the blood



Keywords:

α -Amino Acids

- **Structure**
- **Side chains**
- **Acid-base properties**
- **Synthesis**

Peptides and Proteins

- **Peptide bond**
- **Nomenclature**
- **Level of protein structure**
- **Functions**