

AMINO ACIDS, PEPTIDES & PROTEINS

↳ ≥ 2 Amino acids linked
by a peptide bond

* # of Peptide bonds = # of Amino acids - 1

- e.g.: a Protein contains 120 amino acids,

How many Peptide bonds it has?

- # of Peptide bonds = $120 - 1$
= 119 bonds

AMINO ACIDS

Biochemical Properties

20 standard amino acids (α-amino acids) – proteins
↳ In mammals

Uncommon amino acids in the body (in free form or in combined states) that are not associated with proteins. For example, citrulline and ornithine (disposal of waste nitrogen via the urea cycle)

- * Excretion process:
 1. Respiration: $\text{CO}_2, \text{H}_2\text{O}$
 2. Sweating: H_2O , Salts
 3. Urination: H^+ , Urea, Creatinine

→ One of the ways by which the body gets rid of nitrogen.

→ Source

Precursors to hormones – tyrosine in the formation of thyroid hormones

↳ Standard a.a, contains benzene ring in side chain along with $(-\text{OH})$

Glutamate – as neurotransmitter

↳ acidic a.a along with Aspartate
↳ Contains Carbonyl group $\rightarrow \text{COOH}$

→ Inhibitory neurotransmitter (GABA)

↳ Glu, Asp, GABA

* any a.a containing benzene ring will have a smell (Aromatic a.a), these include:
- Tyrosin
- Phenyl Alanin
- Tryptophan

- Tryptophan has two rings, thus it can absorb & emit light \rightarrow Flourescent

Section 5.1: Amino Acids

* Smallest a.a is Glycine

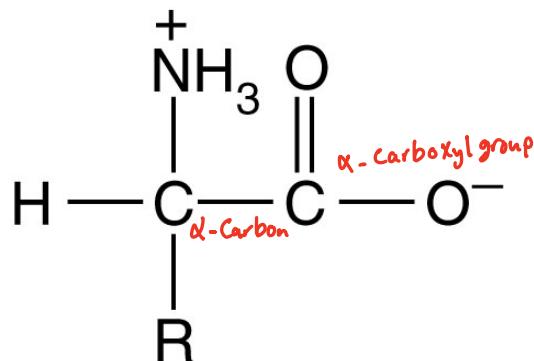


Figure 5.3 General Structure of the α -Amino Acids



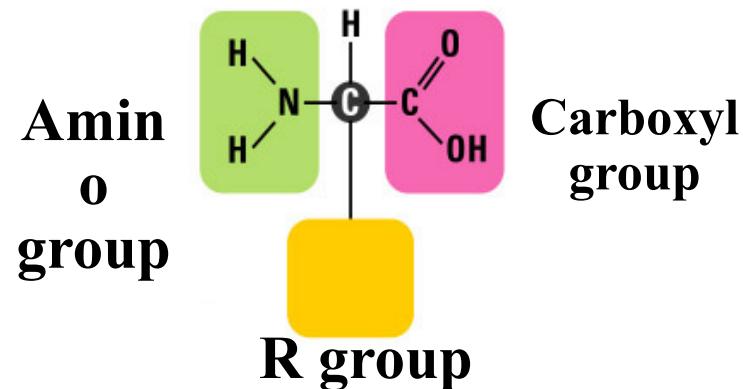
* Basic Amino Acids:

- Histidine
- Lysine
- Arginine

- 19 have the same general structure: central (α) carbon, an amino group, carboxylate group, hydrogen atom, and an R group (proline is the exception)
- At pH 7, the carboxyl group is in its dissociated conjugate base form ($-\text{COO}^-$) while the amino group is its undissociated conjugate acid form ($-\text{NH}_3^+$); therefore, it is amphoteric

Structure of Amino Acids

Amino acids have a central carbon with 4 things bonded to it:



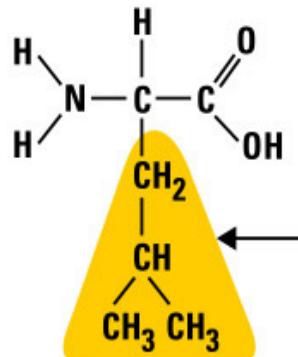
Amino group -NH₃

Carboxyl group -COOH

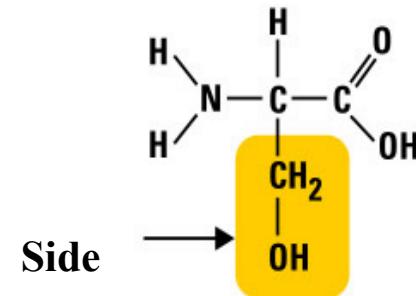
Hydrogen -H

Side group -R

chain



Leucine -hydrophobic



Serine-hydrophilic

- **Amino Acid Stereoisomers** → *2nd diplo class*
- Because the α -carbon (**chiral carbon**) is attached to four different groups, they can exist as **stereoisomers**
- Except glycine, which is the only nonchiral standard amino acid

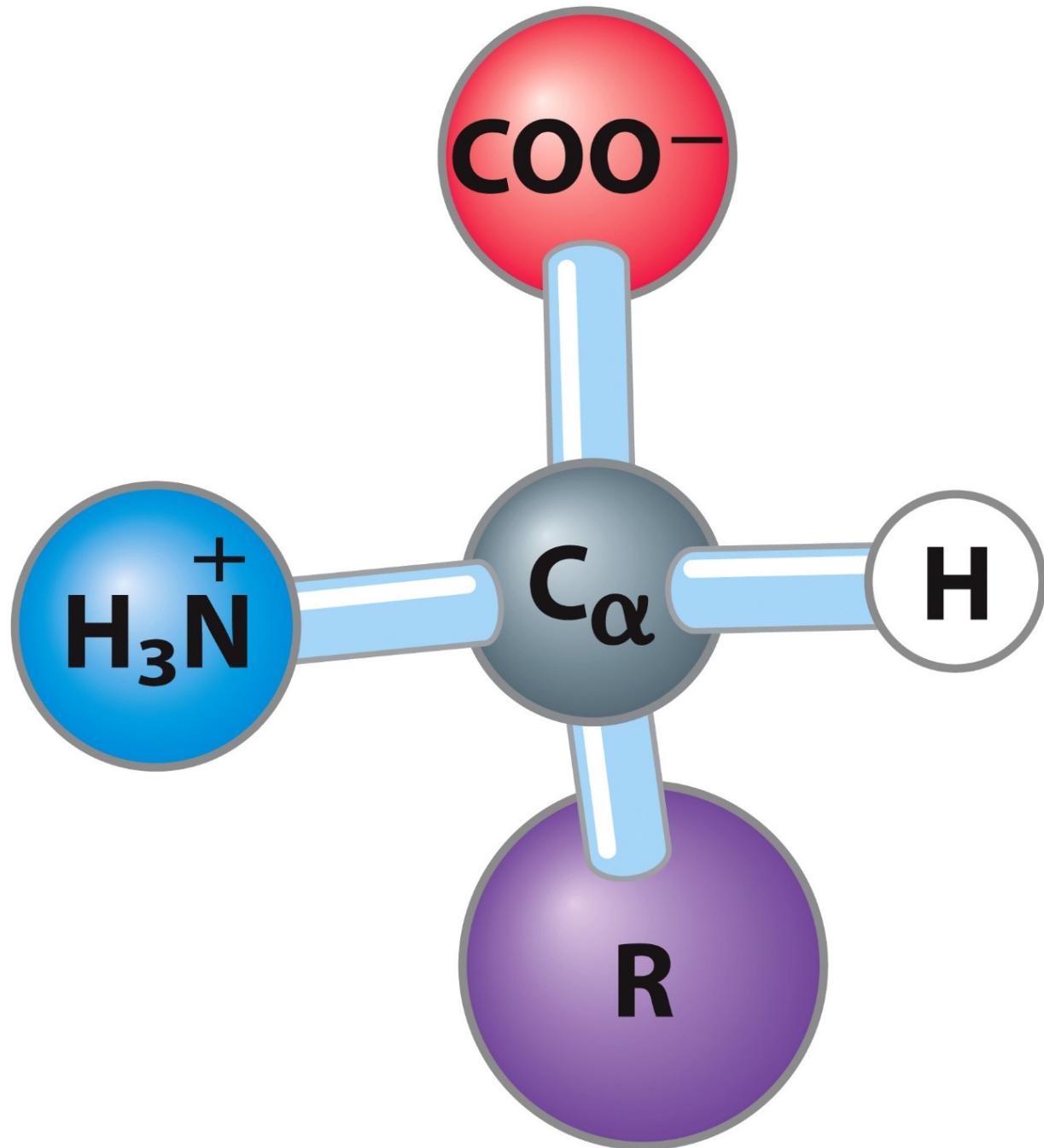
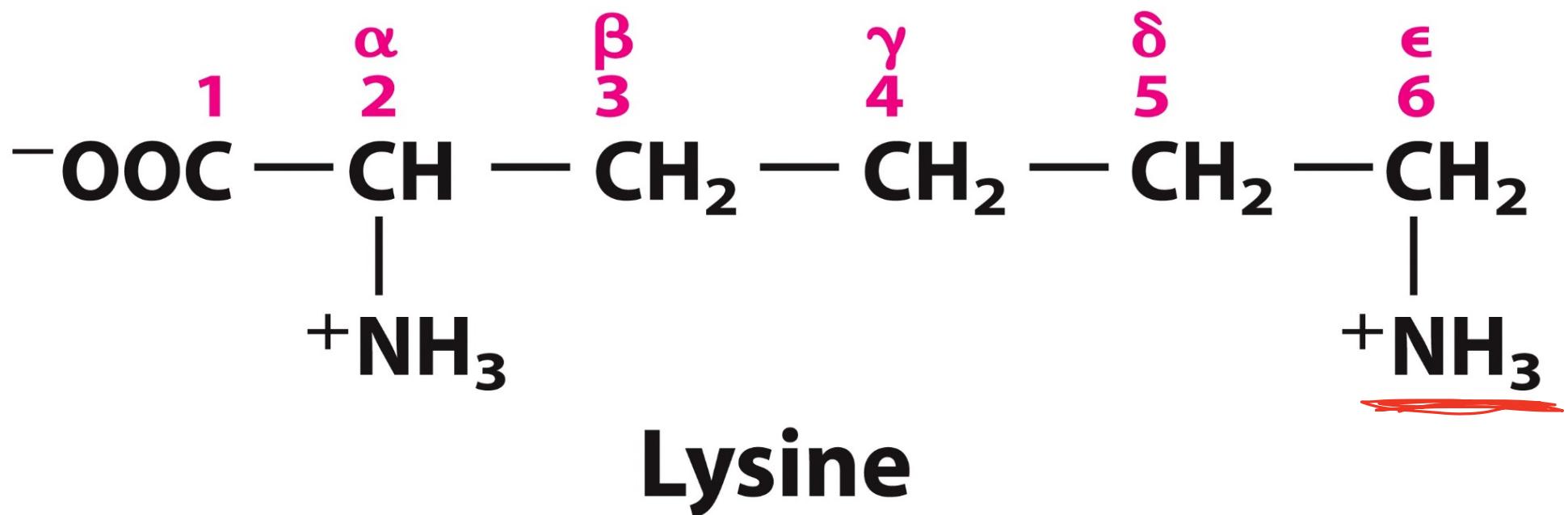


Figure 3-2

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Section 5.1: Amino Acids

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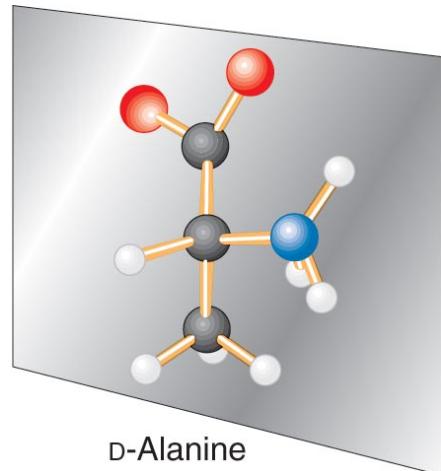


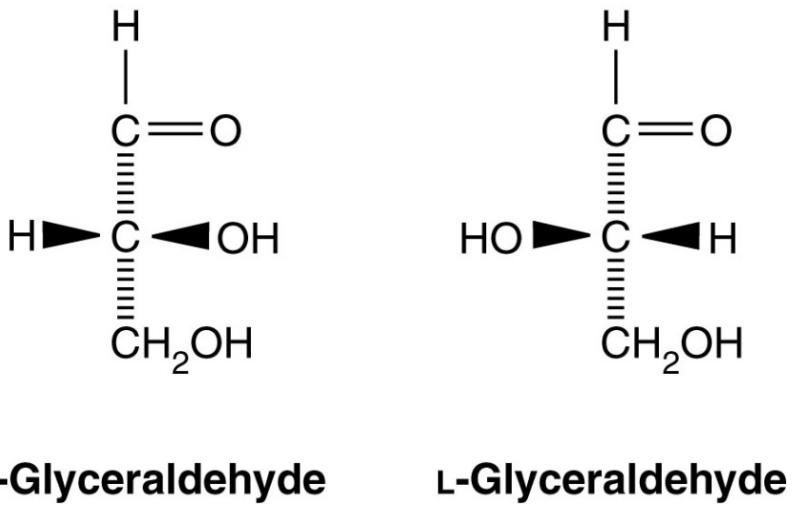
Figure 5.7 Two Enantiomers

- The molecules are mirror images of one another, or **enantiomers**
- They cannot be superimposed over one another and rotate plane, polarized light in opposite directions (**optical isomers**)

Section 5.1: Amino Acids

* just remember that Carbohydrates exist in our bodies in the D form

Figure 5.8 D- and L-Glyceraldehyde



- Molecules are designated as D or L (glyceraldehyde is the reference compound for optical isomers)
- D or L is used to indicate the similarity of the arrangement of atoms around the molecule's asymmetric carbon to the asymmetric carbon of the glyceraldehyde isomers
- Chirality has a profound effect on the structure and function of proteins

Cur
view

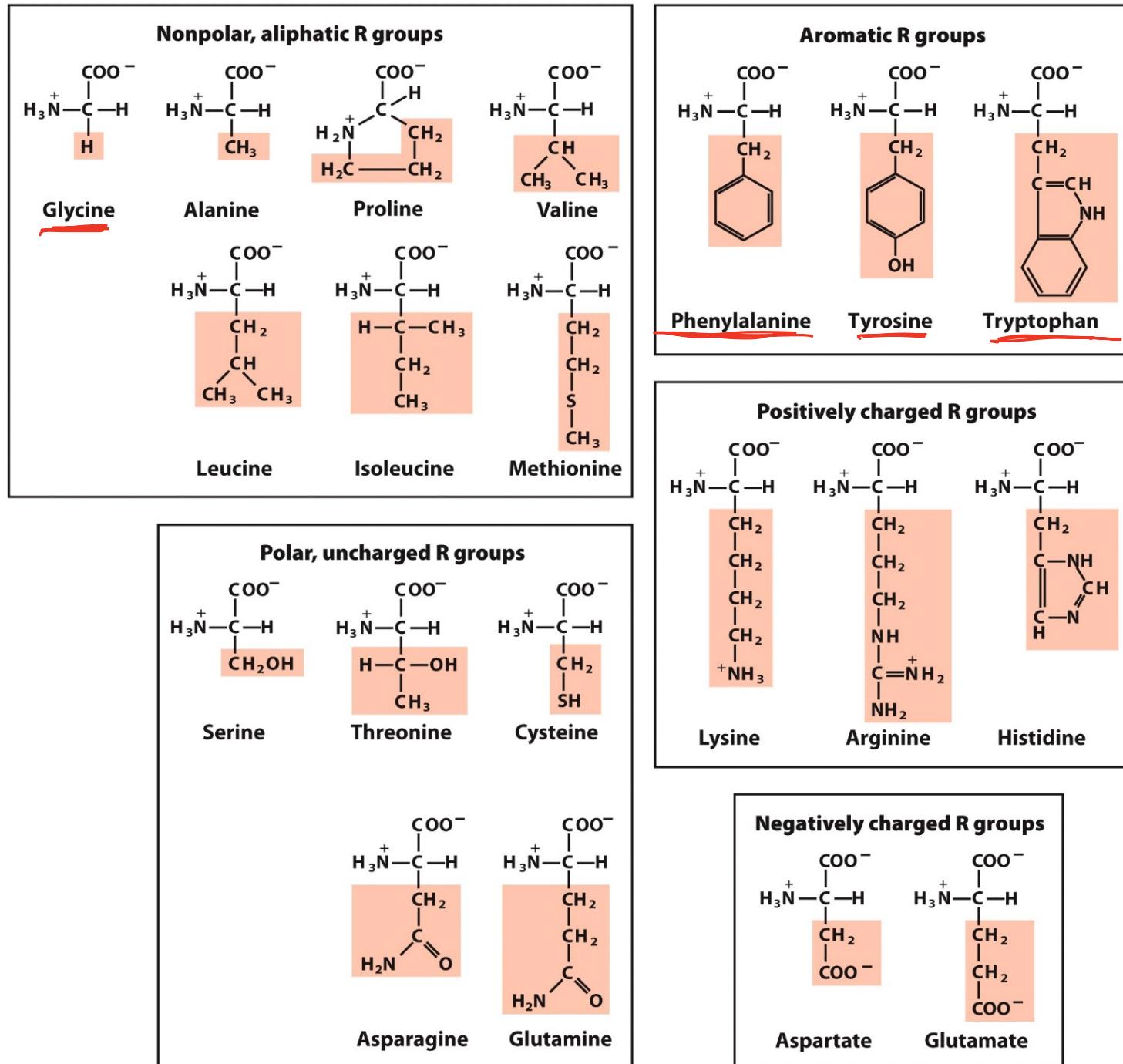


Figure 3-5

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Section 5.1: Amino Acids

TABLE 5.1 Names and Abbreviations of the Standard Amino Acids

| Amino Acid | Three-Letter Abbreviation | One-Letter Abbreviation |
|---------------|---------------------------|-------------------------|
| Alanine | Ala | A |
| Arginine | Arg | R |
| Asparagine | Asn | N |
| Aspartic acid | Asp | D |
| Cysteine | Cys | C |
| Glutamic acid | Glu | E |
| Glutamine | Gln | Q |
| Glycine | Gly | G |
| Histidine | His | H |
| Isoleucine | Ile | I |
| Leucine | Leu | L |
| Lysine | Lys | K |
| Methionine | Met | M |
| Phenylalanine | Phe | F |
| Proline | Pro | P |
| Serine | Ser | S |
| Threonine | Thr | T |
| Tryptophan | Trp | W |
| Tyrosine | Tyr | Y |
| Valine | Val | V |

Section 5.1: Amino Acids

• Biologically Active Amino Acids

- Amino acids can have other biological roles

1. Some amino acids or derivatives can act as chemical messengers

- Neurotransmitters (tryptophan- derivative serotonin) and hormones (tyrosine-derivative thyroxine)

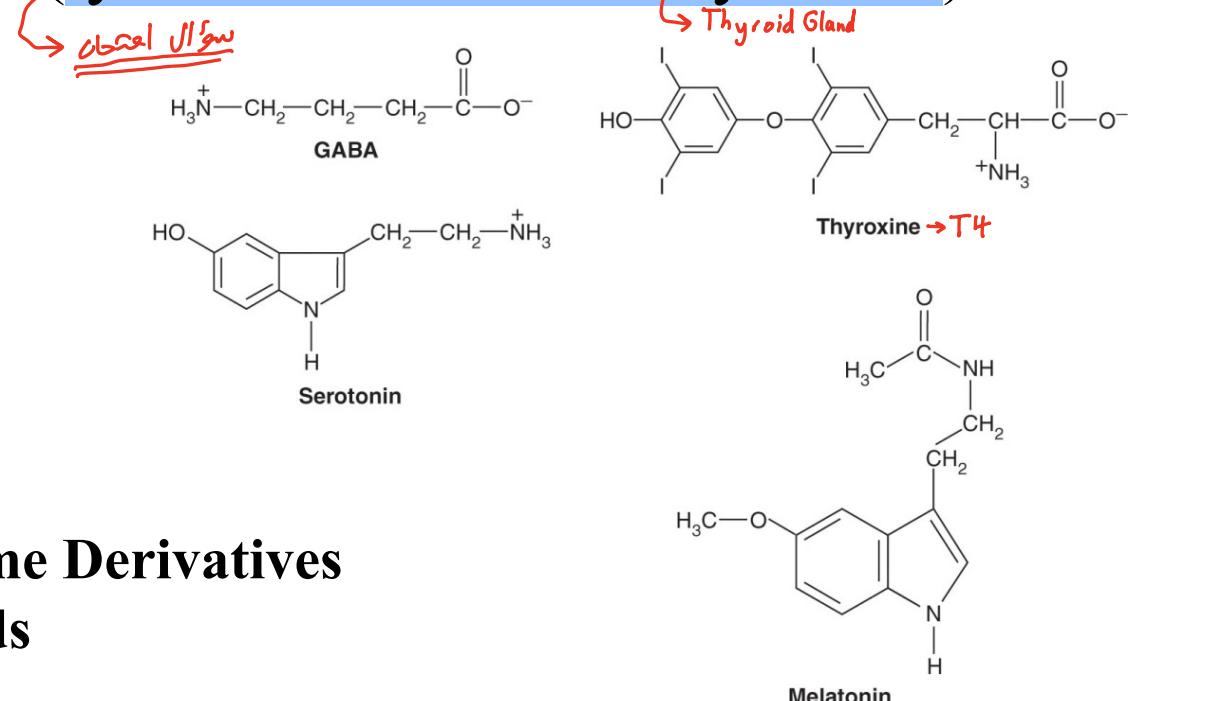


Figure 5.4 Some Derivatives of Amino Acids

Section 5.1: Amino Acids

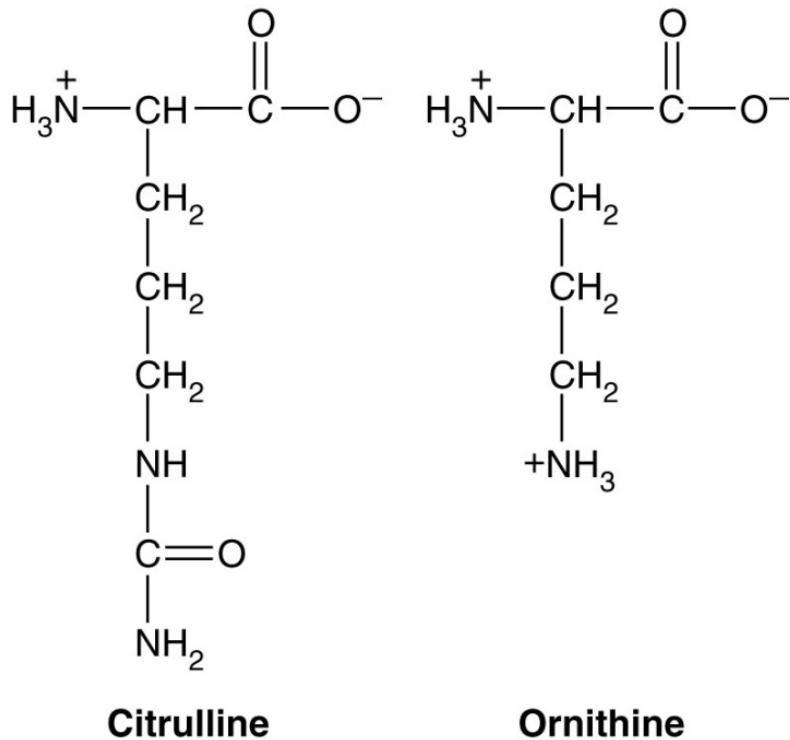


Figure 5.5 Citruline and Ornithine

* Heme: **Tetrapyrrol**
 - important for O_2 binding with a.a. sidechains



or other **Heme Proteins** \rightarrow "Super Family"
 s and heme) \downarrow Hundreds of Protein

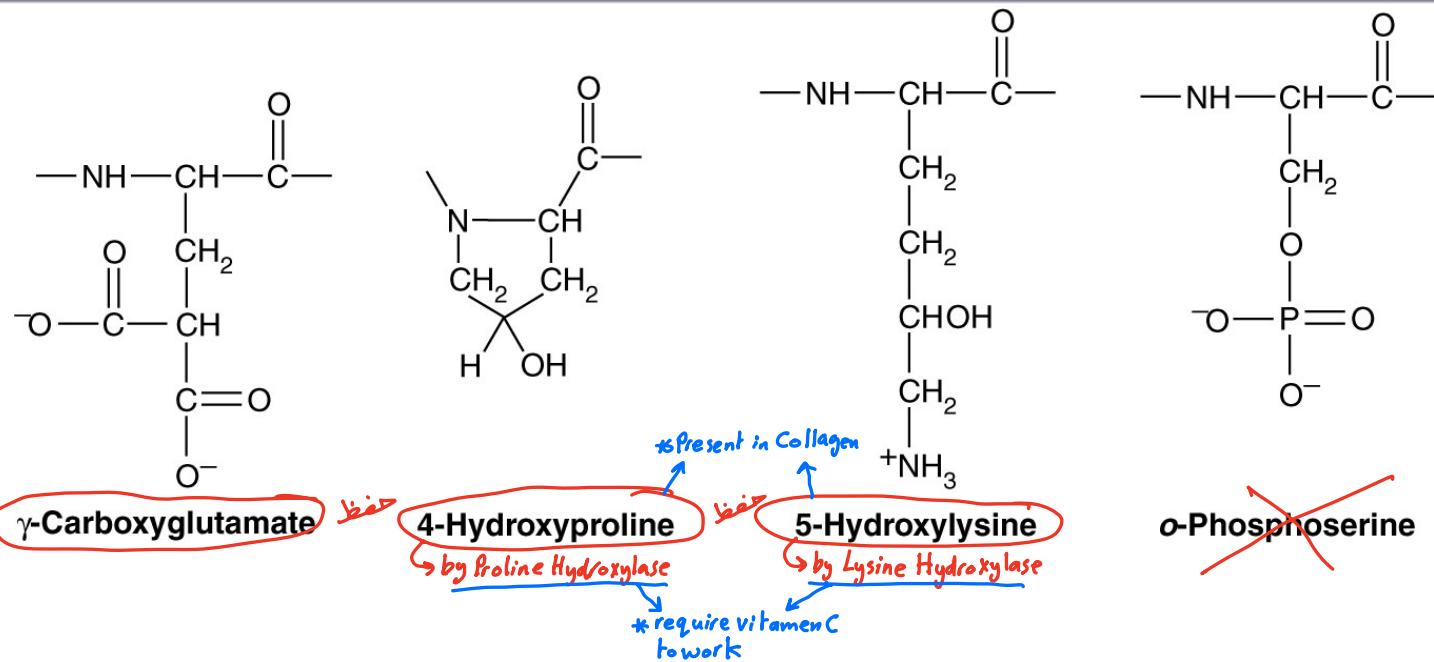
e.g.: **Cytochromes** \hookrightarrow Produced in the liver and deal with hormones and drugs

2. Act as precursors for other molecules (nucleotides and heme)
 - ↳ Nucleic acids
 - ↳ Heme Proteins, "Super Family"
e.g.: Cytochromes
↳ Produced in the liver and deal with hormones and drugs
Hundreds of Proteins
3. Metabolic intermediates (arginine, ornithine, and citrulline in the urea cycle) ↳ Essential A.As: A.As which Can't be produced

- Essential A.A.s: A.A.s which can't be produced inside the body therefore they must be obtained from diet.
 - Arginine is essential during infancy & childhood, because hepatocytes are not mature enough to produce required enzymes.

Section 5.1: Amino Acids

Figure 5.6 Modified Amino Acid Residues Found in Polypeptides



• Modified Amino Acids in Proteins

- Some proteins have amino acids that are modified after synthesis
 - Serine, threonine, and tyrosine can be phosphorylated
 - γ -Carboxyglutamate (prothrombin), 4-hydroxyproline (collagen), and 5-hydroxylysine (collagen)

structural Proteins don't dissolve in water
Elastin
most abundant Protein in the body

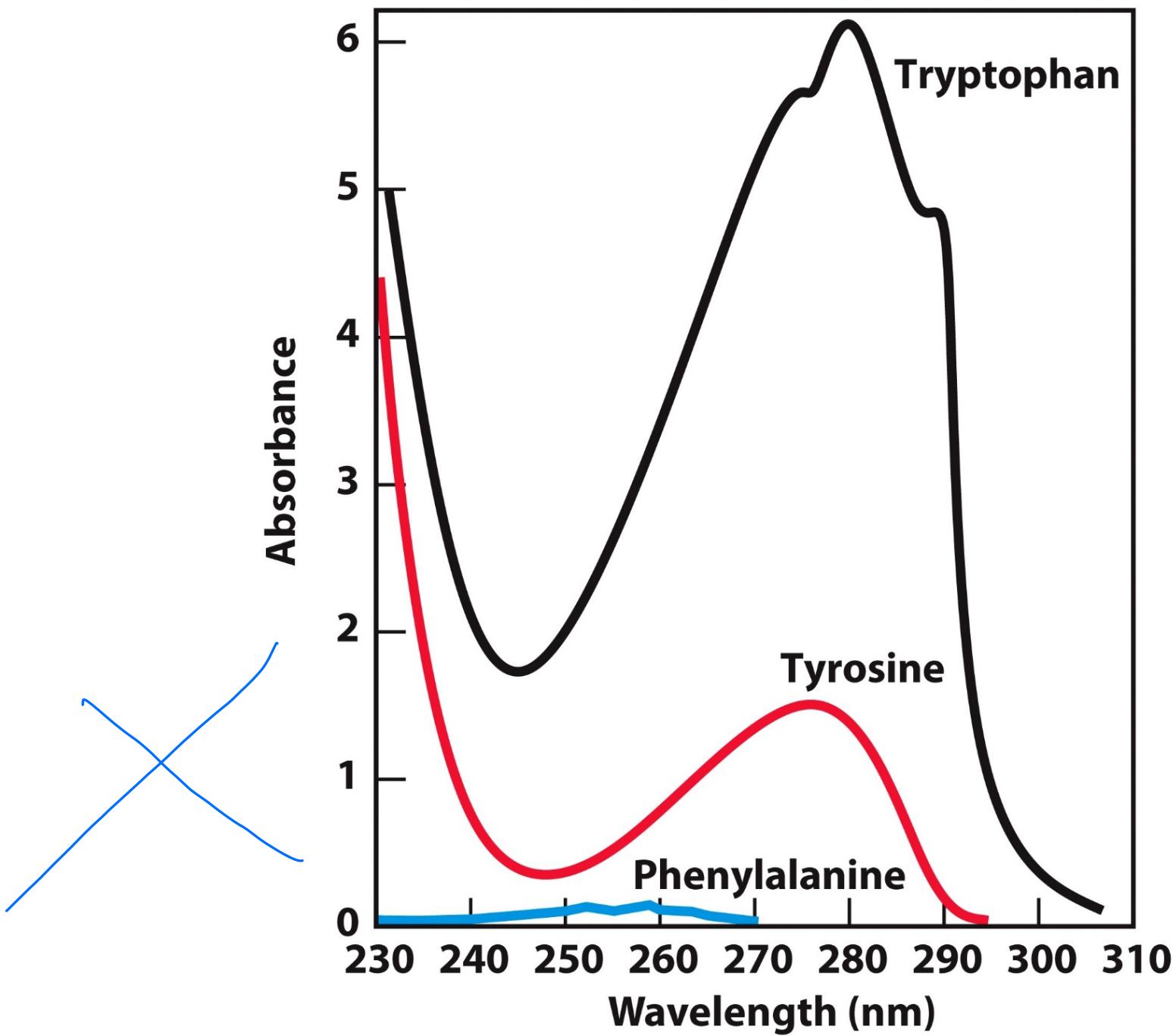
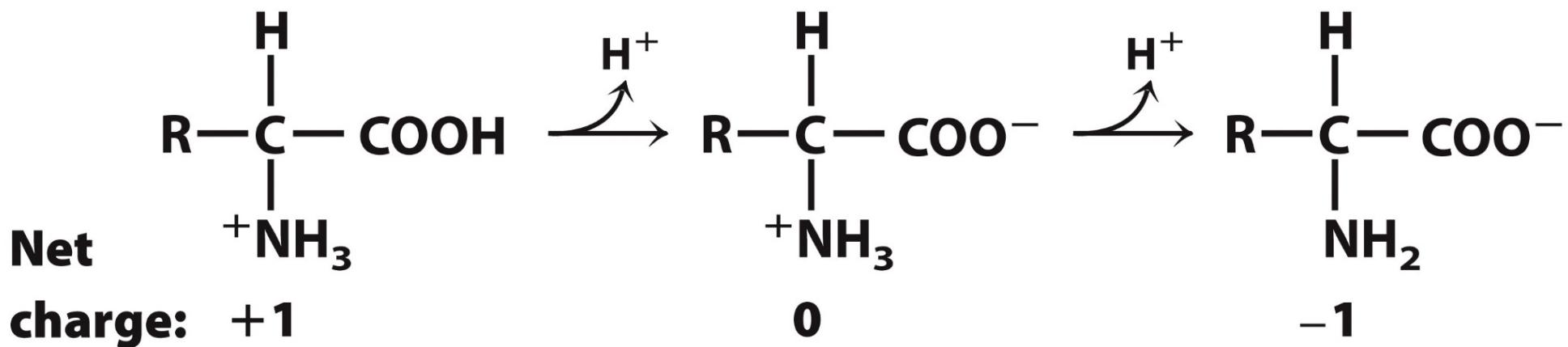


Figure 3-6

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- Molecules that have both positive and negative charges on different atoms are **zwitterions** and have no net charge at pH 7
 For Non-polar A.A.s
- The R group is what gives the amino acid its unique properties



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Acid-Base Properties of Amino Acids

The α -COOH and α -NH₂ groups in amino acids are capable of ionizing (as are the acidic and basic R groups of the amino acids)

الحالة: $-\text{COO}^-$: unprotonated/dissociated

At physiological pH (around 7.4), the carboxyl group will be unprotonated^{dissociated} and the amino group will be protonated. An amino acid with no ionizable R group would be electrically neutral at this pH. This species is termed as a zwitterion

When the net charge of an amino acid or protein is zero, the pH will be equivalent to the isoelectric point (pI)

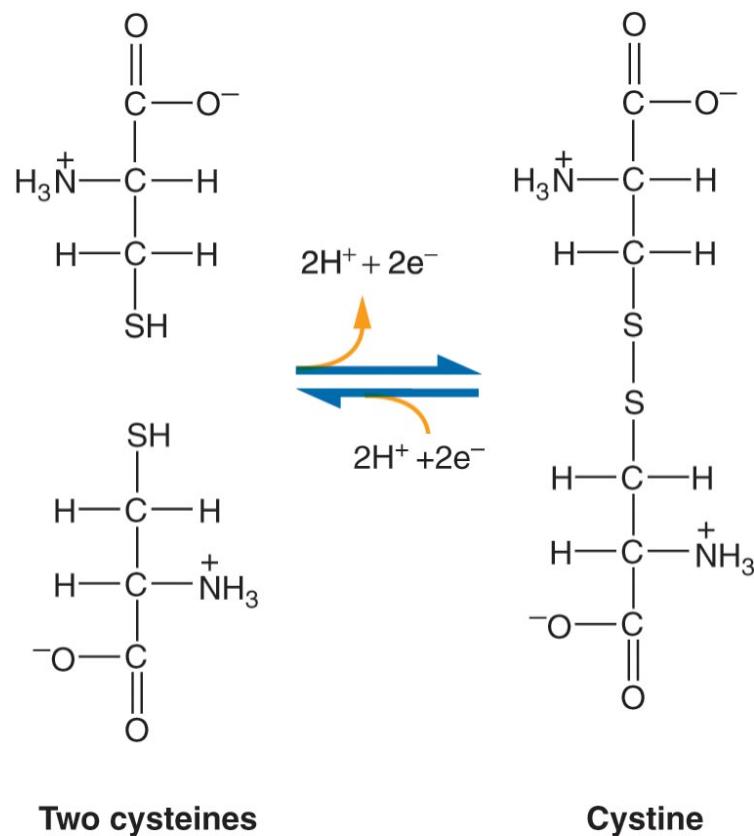
Histidine

The imidazole ring of histidine allows it to act as either a proton donor or acceptor at physiological pH

Frequently found in the reactive center of enzymes
binding site/Active site

Histidine residues in hemoglobin are **critical** in its ability
~~X~~ to buffer the H⁺ from carbonic acid ionization in red blood cells (RBCs)

Section 5.1: Amino Acids



- **Cysteine oxidation** leads to a reversible disulfide bond
 - A **disulfide bridge** forms when two cysteine residues form this bond
 - Helps stabilize polypeptides and proteins

Figure 5.12 Oxidation of Two Cysteine Molecules to Form Cystine



PEPTIDES

Peptides

Peptides – <30 amino acids

Proteins - >100 amino acids joined by peptide bonds

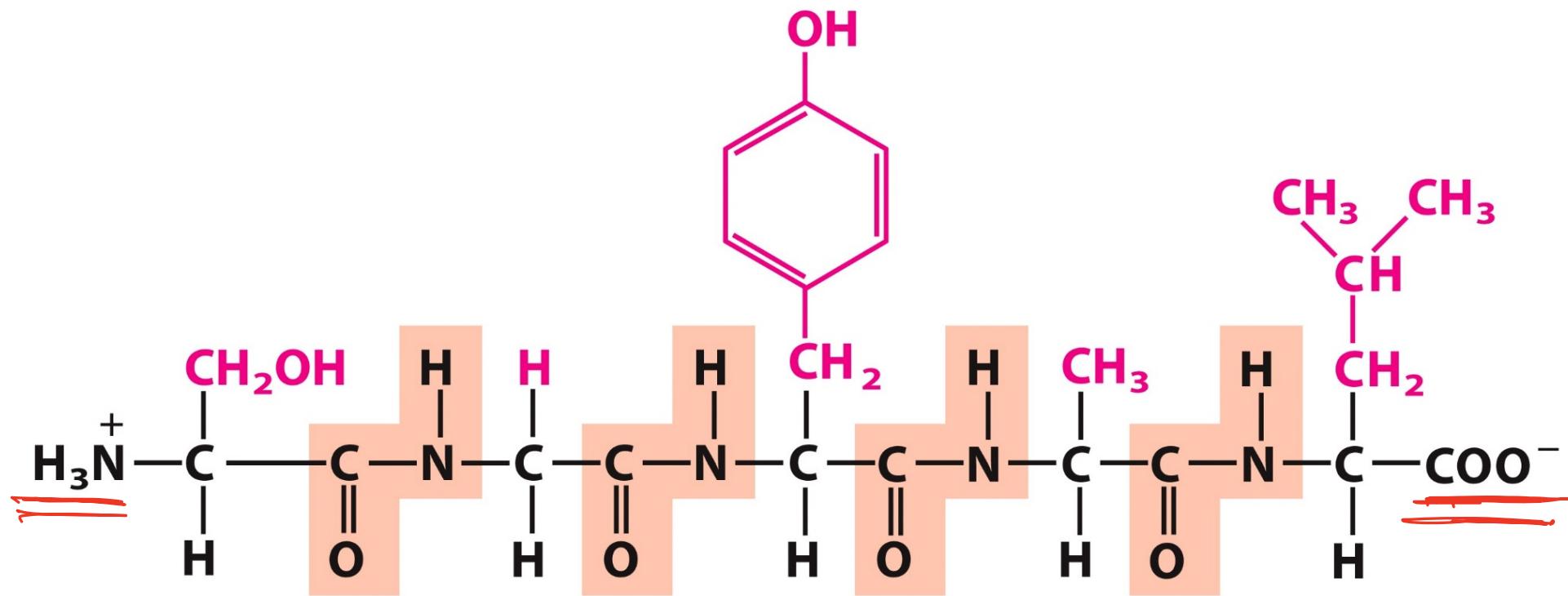
Polypeptide chain – the side chains and backbone groups interact with each other through many weak interactions, including van der Waals, hydrogen bonds, electrostatic interactions, as well as hydrophobic effect (non-polar groups) to bring about a protein's shape and target interactions

→ Non-covalent

Section 5.2: Peptides

cycle line

- ~~Less structurally complex than larger proteins, peptides still have biologically important functions~~
- **Glutathione** is a tripeptide found in most all organisms and is involved in protein and DNA synthesis, toxic substance metabolism, and amino acid transport
- **Vasopressin** is an antidiuretic hormone that regulates water balance, appetite, and body temperature
- **Oxytocin** is a peptide that aids in uterine contraction and lactation



**Amino-
N-terminal end**

**Carboxyl-
C-terminal end**

* In any Protein, we have
only 1 α -aminogroup &
1 α -Carboxyl group

Figure 3-14
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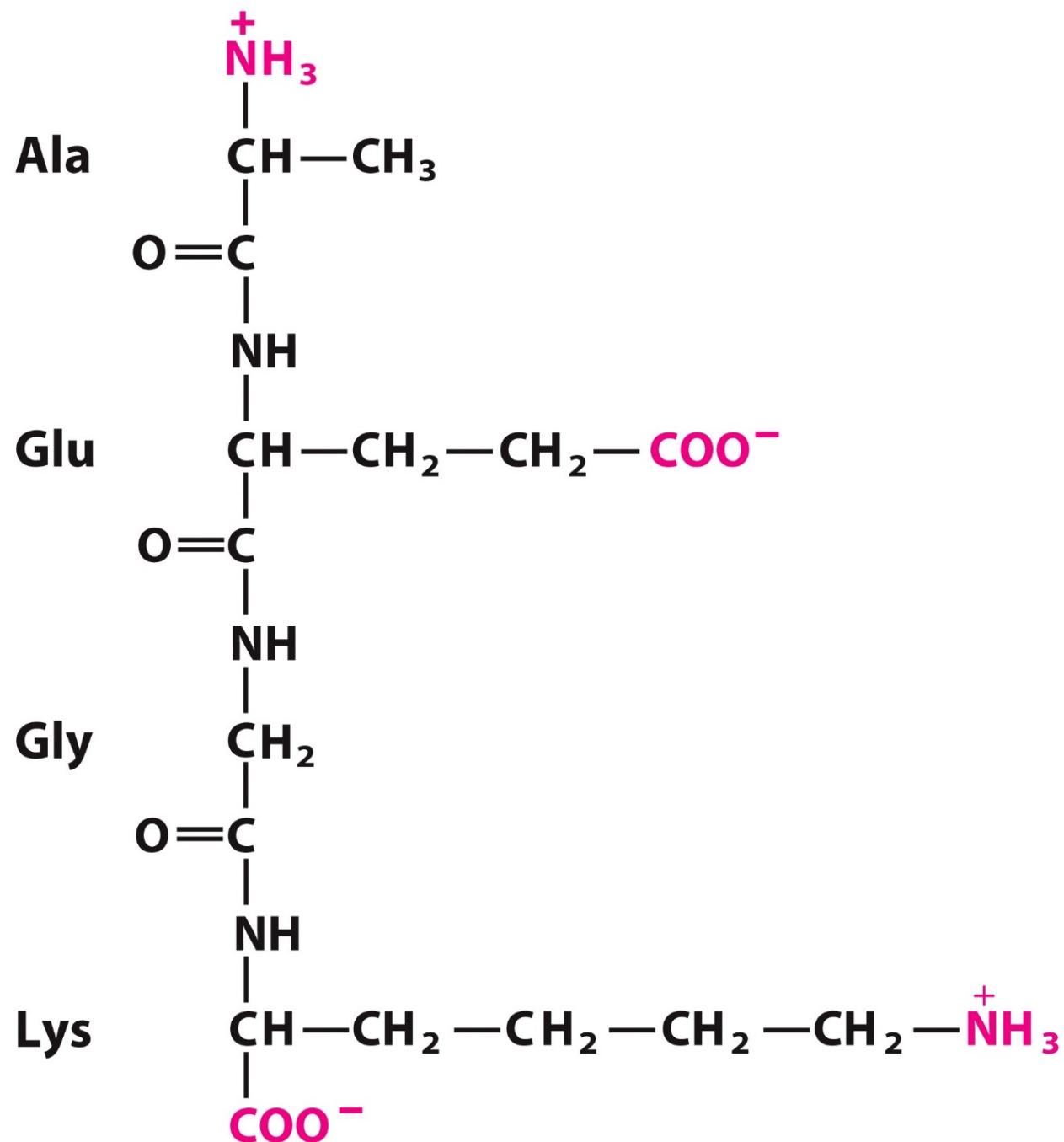


Figure 3-15

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Figure 3-15

This tetrapeptide (Alanyl glutamyl glycyl lysine) has one free α -amino group, one free α -carboxyl group, and two ionizable R groups. The groups ionized at pH 7.0 are in red. Ionizable R groups in a peptide also contribute to the overall acid-base properties of the molecule

Biologically Active Peptides

Naturally occurring peptides: 2 – 1000 amino acids

The artificial sweetener Aspartame (also called Nutra Sweet) – commercially synthesized dipeptide L-aspartyl-L-phenylalanyl methyl ester

Snake venoms (20 – 100 components) – the majority of these are peptides and proteins

α -amanitin (small peptide) – extremely toxic poison from the mushroom amanita phalloides

PROTEINS

TABLE 3-2 Molecular Data on Some Proteins

| | Molecular weight | Number of residues | Number of polypeptide chains |
|--|-------------------------|---------------------------|-------------------------------------|
| Cytochrome c (human) | 12,400 | 104 | 1 |
| Ribonuclease A (bovine pancreas) | 13,700 | 124 | 1 |
| Lysozyme (chicken egg white) | 14,300 | 129 | 1 |
| Myoglobin (equine heart) | 16,700 | 153 | 1 |
| Chymotrypsin (bovine pancreas) | 25,200 | 241 | 3 |
| Chymotrypsinogen (bovine) | 25,700 | 245 | 1 |
| Hemoglobin (human) | 64,500 | 574 | 4 |
| Serum albumin (human) | 66,000 | 609 | 1 |
| Hexokinase (yeast) | 107,900 | 972 | 2 |
| RNA polymerase (<i>E. coli</i>) | 450,000 | 4,158 | 5 |
| Apolipoprotein B (human) | 513,000 | 4,536 | 1 |
| Glutamine synthetase (<i>E. coli</i>) | 619,000 | 5,628 | 12 |
| Titin (human) | 2,993,000 | 26,926 | 1 |

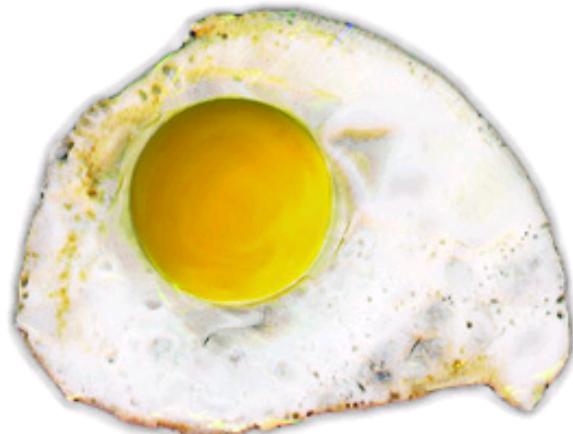
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Changes in temperature & pH can denature (unfold) a protein so it no longer works

Denaturating Proteins

Cooking denatures protein in eggs



Milk protein separates into curds & whey when it denatures

- **DENATURING CONDITIONS**

1. Strong acid or base
2. Organic solvents
3. Detergents
4. Reducing agents
5. Salt concentration
6. Heavy metal ions
7. Temperature

