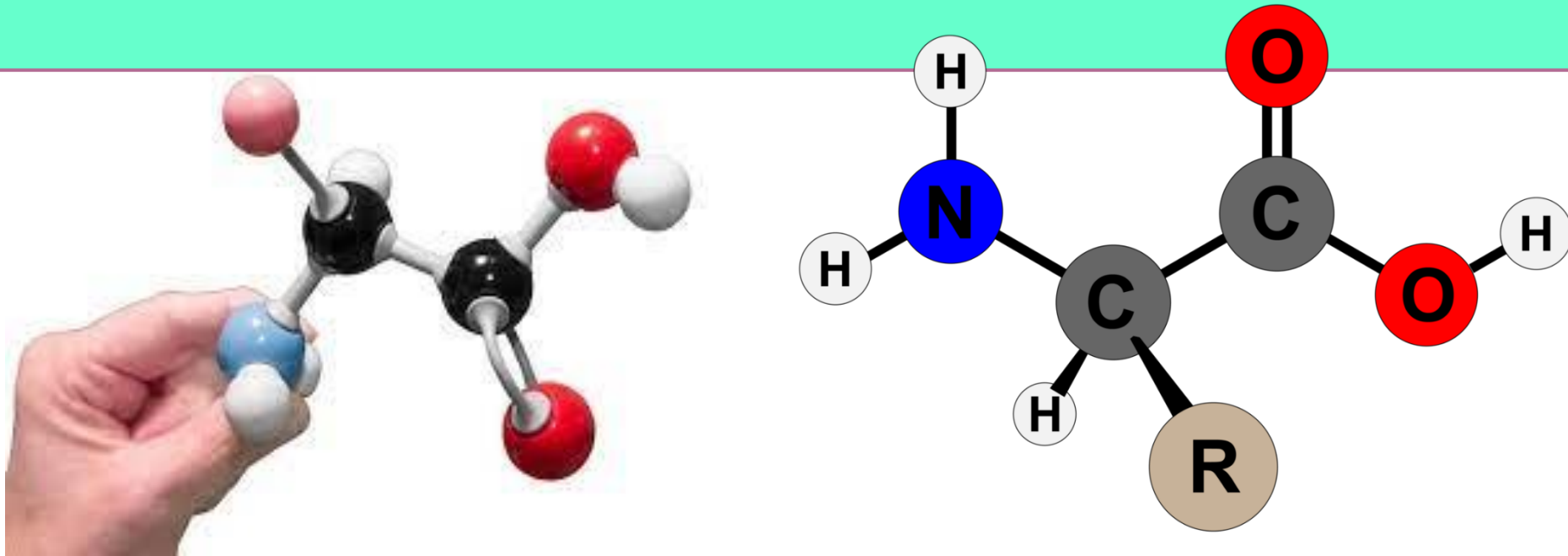


CHAPTER 12

AMINO ACIDS

12.1 INTRODUCTION TO AMINO ACIDS
12.2 CHEMICAL PROPERTIES

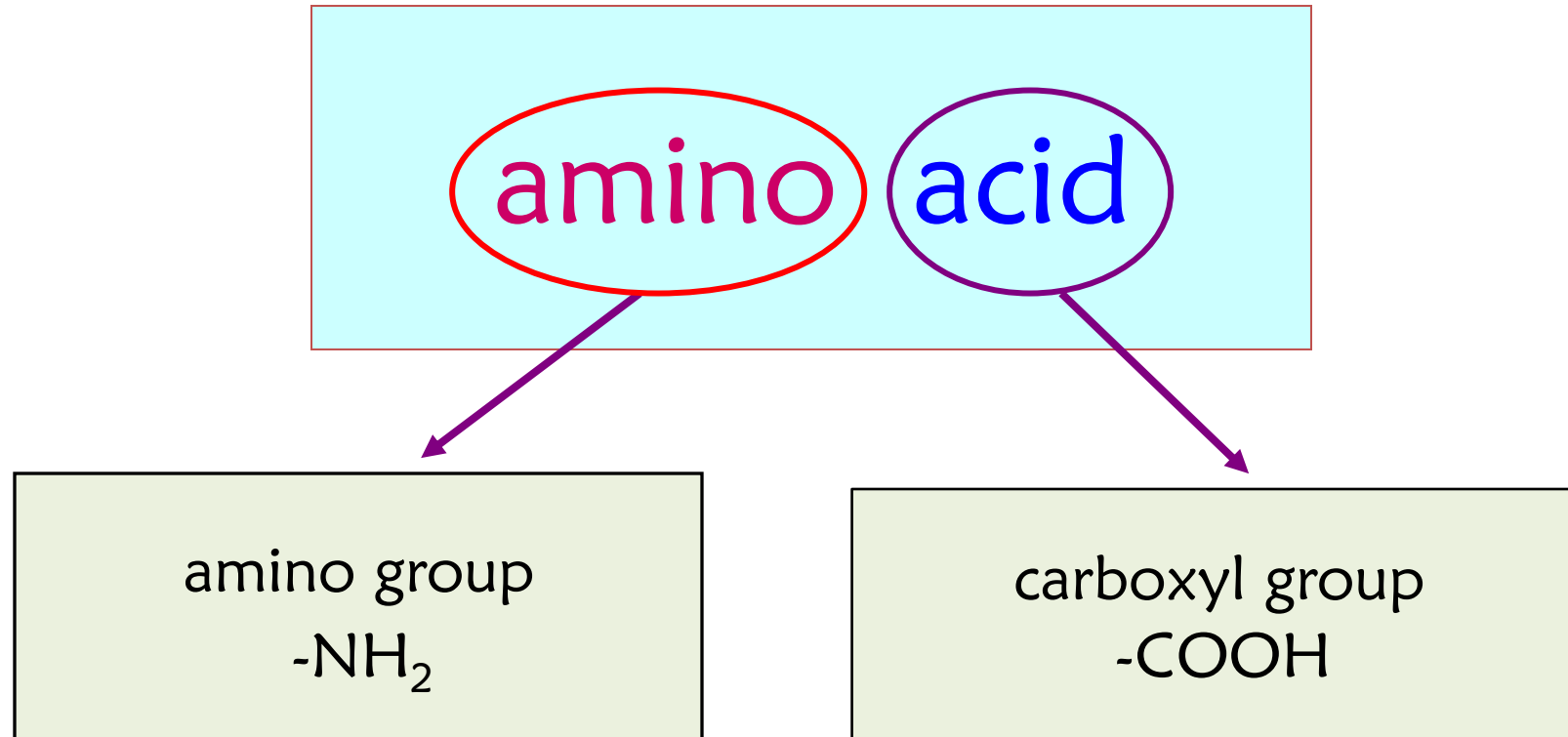


12.1 : INTRODUCTION TO AMINO ACIDS

Students should be able to:

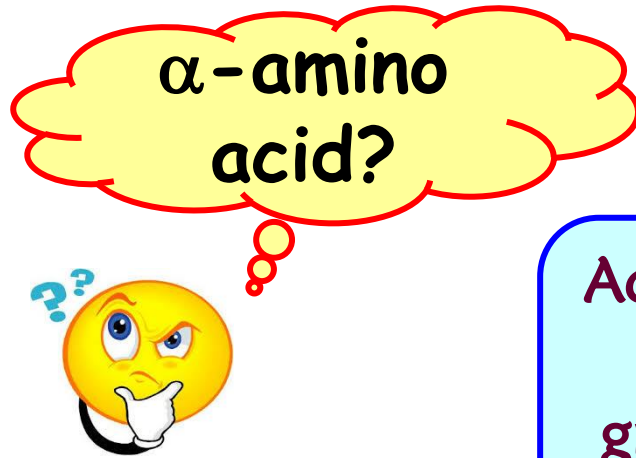
- a) Give the name for a given amino acid according to the IUPAC nomenclature (C2)
- b) Give the general structure of α -amino acid in 12.1(a). (C2)
- c) Define the terms: (C1)
 - i. zwitterion;
 - ii. Isoelectric point (pI).
- d) Predict the structure of a given amino acid, (C3)
 - i. in acidic medium
 - ii. in basic medium
 - iii. at pI.

12.1 : INTRODUCTION TO AMINO ACIDS



Amino acids are molecules with two functional groups:
amino group (-NH₂) and **carboxyl group (-COOH)**

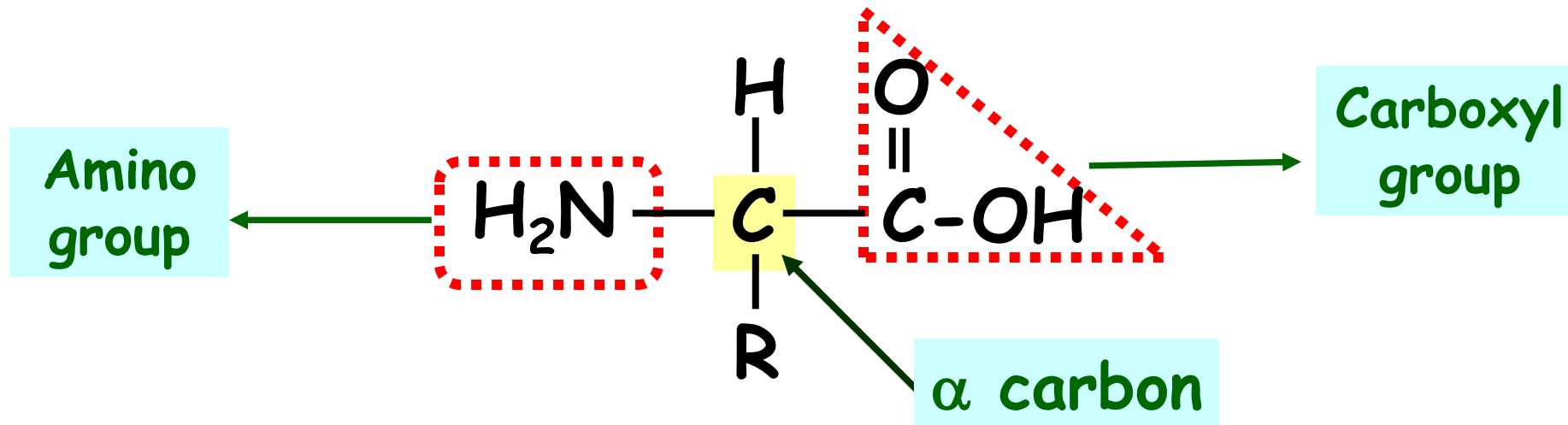
Amino acid commonly found in proteins are α -amino acid



Acid amino on which both carboxyl and amino groups are bonded to the **SAME CARBON** atom.



General structure of α -amino acid

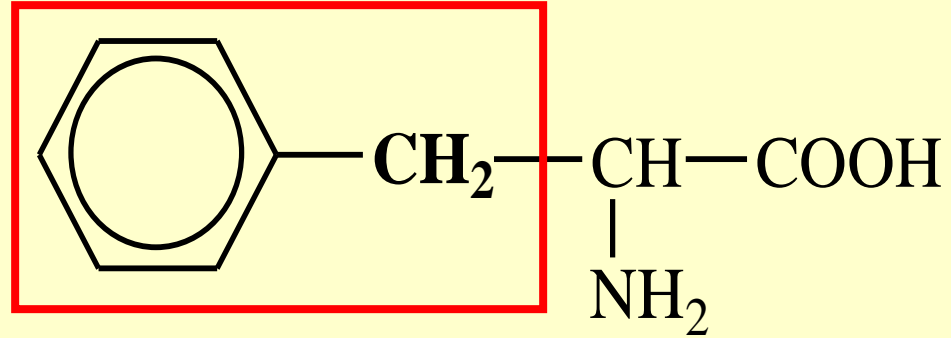


Classification of 20 amino acids in protein.

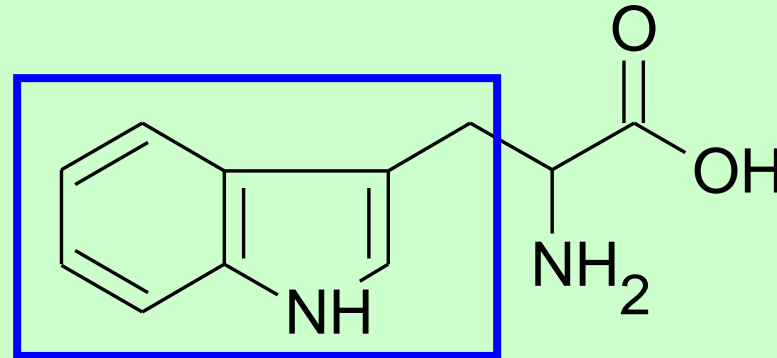
Name	Structure
Glycine (Gly)	$\text{H} - \text{CH} - \text{COOH}$ \mid NH_2
Alanine (Ala)	$\text{H}_3\text{C} - \text{CH} - \text{COOH}$ \mid NH_2

Valine (Val)	$ \begin{array}{c} \text{CH}_3 \\ \\ \text{HC} - \text{CH} - \text{COOH} \\ \quad \\ \text{CH}_3 \quad \text{NH}_2 \end{array} $
Leucine (Leu)	$ \begin{array}{c} \text{CH}_3 \\ \\ \text{CHCH}_2 - \text{CH} - \text{COOH} \\ \quad \\ \text{CH}_3 \quad \text{NH}_2 \end{array} $
Isoleucine (Ile)	$ \begin{array}{c} \text{CH}_3\text{CH}_2\text{CH} - \text{CH} - \text{COOH} \\ \quad \\ \text{CH}_3 \quad \text{NH}_2 \end{array} $

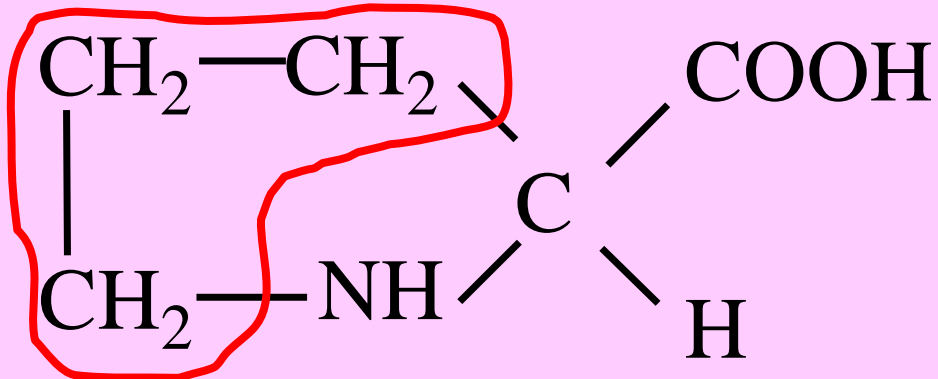
Phenylalanine
(Phe)



Tryptophan
(Trp)



Proline
(Pro)



<p>Serine (Ser)</p>	$\boxed{\text{HO}-\text{CH}_2}-\underset{\substack{ \\ \text{NH}_2}}{\text{CH}}-\text{COOH}$
<p>Threonine (Thr)</p>	$\begin{array}{c} \text{OH} \\ \\ \text{CH}_3-\text{C}-\underset{\substack{ \\ \text{NH}_2}}{\text{CH}}-\text{COOH} \\ \\ \text{H} \end{array}$
<p>Cysteine (Cys)</p>	$\text{HS}-\text{CH}_2-\underset{\substack{ \\ \text{NH}_2}}{\text{CH}}-\text{COOH}$

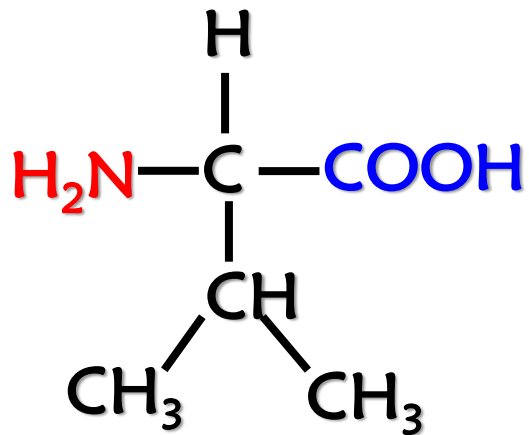
<p>Methionine (Met)</p>	$\text{CH}_3-\text{S}-\text{CH}_2\text{CH}_2-\underset{\text{NH}_2}{\text{CH}}-\text{COOH}$
<p>Tyrosine (Tyr)</p>	$\text{HO}-\text{C}_6\text{H}_4-\text{CH}_2-\underset{\text{NH}_2}{\text{CH}}-\text{COOH}$
<p>Asparagine (Asn)</p>	$\text{NH}_2-\underset{\text{O}}{\underset{\parallel}{\text{C}}}-\text{CH}_2-\underset{\text{NH}_2}{\text{CH}}-\text{COOH}$

Glutamine (Gln)	$\text{NH}_2 - \underset{\text{O}}{\underset{\parallel}{\text{C}}} - \text{CH}_2\text{CH}_2 - \underset{\text{NH}_2}{\underset{ }{\text{CH}}} - \text{COOH}$
Aspartic acid (Asp)	$\text{HOOC} - \text{CH}_2 - \underset{\text{NH}_2}{\underset{ }{\text{CH}}} - \text{COOH}$
Glutamic acid (Glu)	$\text{HOOC} - \text{CH}_2\text{CH}_2 - \underset{\text{NH}_2}{\underset{ }{\text{CH}}} - \text{COOH}$

Lysine (Lys)	$\text{H}_2\text{N}-\text{CH}_2\text{CH}_2\text{CH}_2\text{CH}_2-\underset{\text{NH}_2}{\text{CH}}-\text{COOH}$
Arginine (Arg)	$\text{H}_2\text{N}-\underset{\text{NH}}{\underset{ }{\text{C}}}-\text{NH}-\text{CH}_2\text{CH}_2\text{CH}_2-\underset{\text{NH}_2}{\text{CH}}-\text{COOH}$
Histidine (His)	$\text{HC}=\underset{\text{N}}{\underset{ }{\text{C}}}-\underset{\text{NH}}{\underset{ }{\text{C}}}-\text{CH}_2-\underset{\text{NH}_2}{\text{CH}}-\text{COOH}$

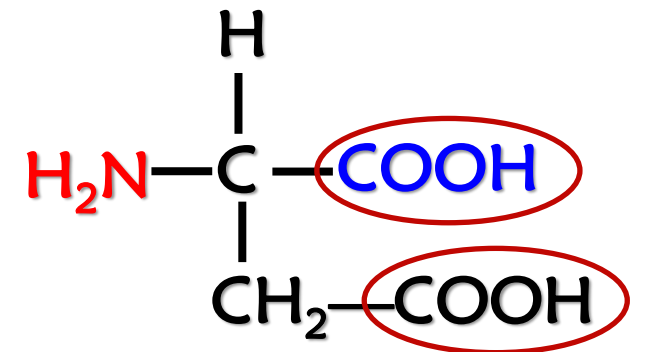
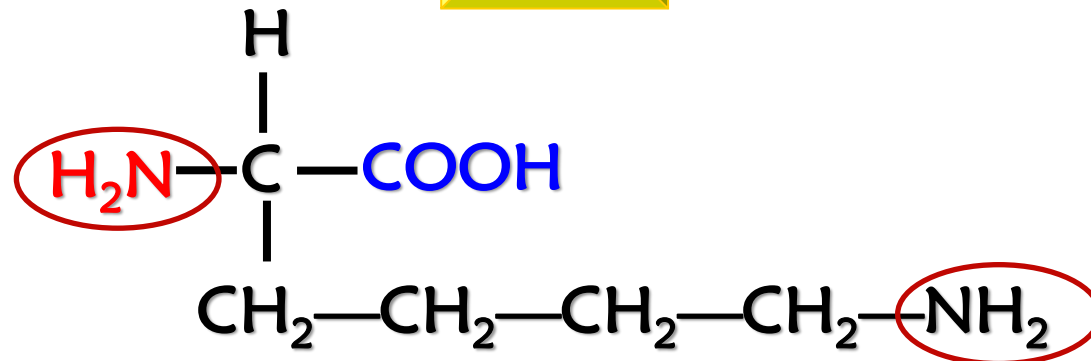
KEEP IN MIND

- ❖ Amino acids with an **additional COOH** group in the side chain - **acidic amino acids**
- ❖ Amino acids with an **additional N** atom in the side chain - **basic amino acids**
- ❖ Others are **neutral amino acids**



neutral

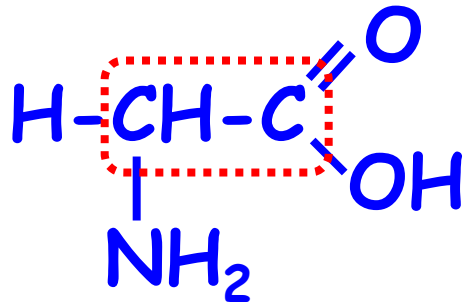
basic



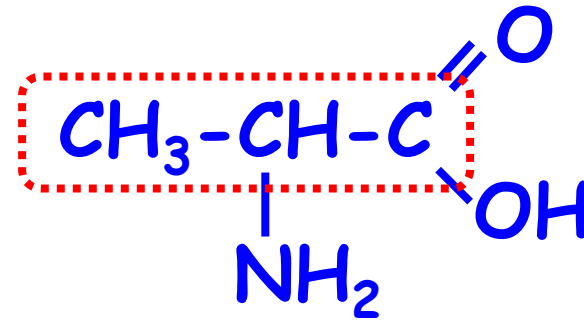
acidic

Nomenclature

- ❑ The names of α -amino acids given are common names.
- ❑ In IUPAC nomenclature, α -amino acids are named as **carboxyl group as the parent name** whereas **amino group as substituent group**.
- ❑ However, IUPAC names are not normally used for α -amino acids



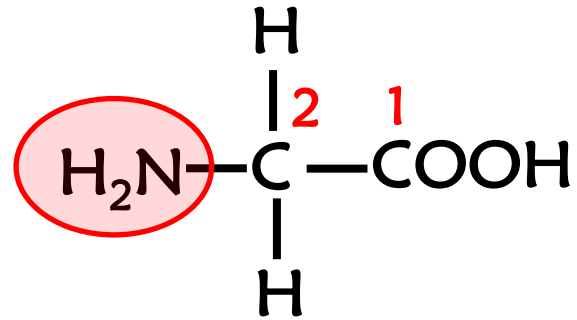
aminoethanoic acid
(glycine)



2-aminopropanoic acid
(alanine)

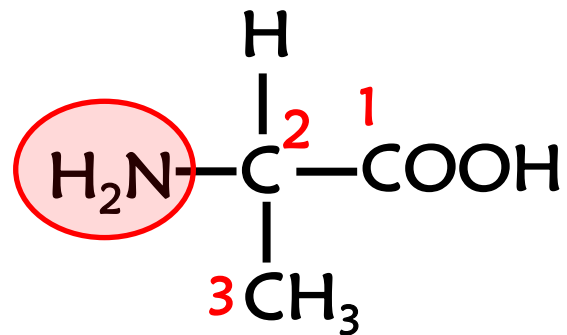
HOW TO NAME THESE AMINO ACID BASED ON IUPAC NOMENCLATURE?

a) glycine



aminoacetic acid / aminoethanoic acid

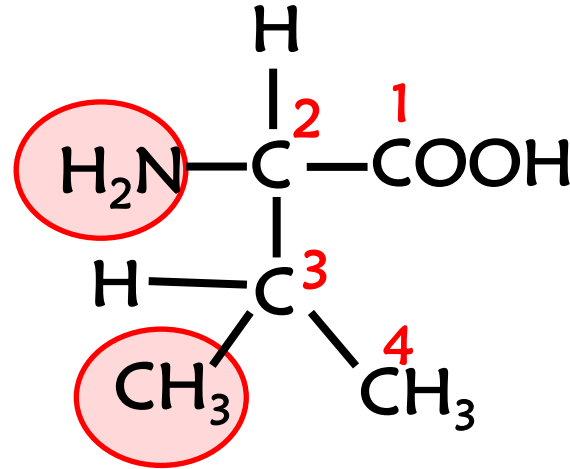
b) alanine



2-aminopropanoic acid

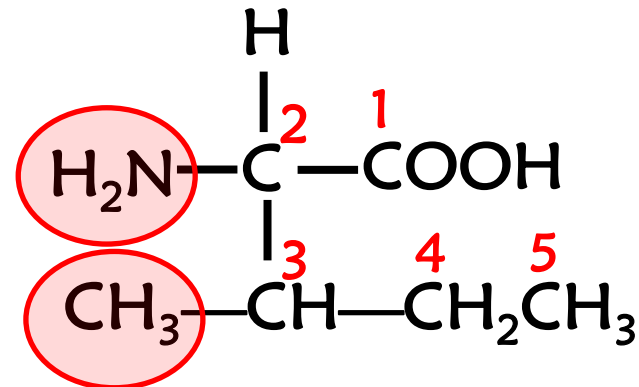
HOW TO NAME THESE AMINO ACID BASED ON IUPAC NOMENCLATURE?

c) valine



2-amino-3-methylbutanoic acid

d) isoleucine



2-amino-3-methylpentanoic acid

Physical Properties

- ❑ Amino acids are white crystalline solids with high melting points.
- ❑ Amino acids such as glycine and alanine dissolve in water to form neutral solution but have low solubility in organic solvents.



This unique properties :

➤ High melting points

➤ High solubility in water (*but low solubility in organic solvents*)

indicates that **amino acids exists as polar ions**.

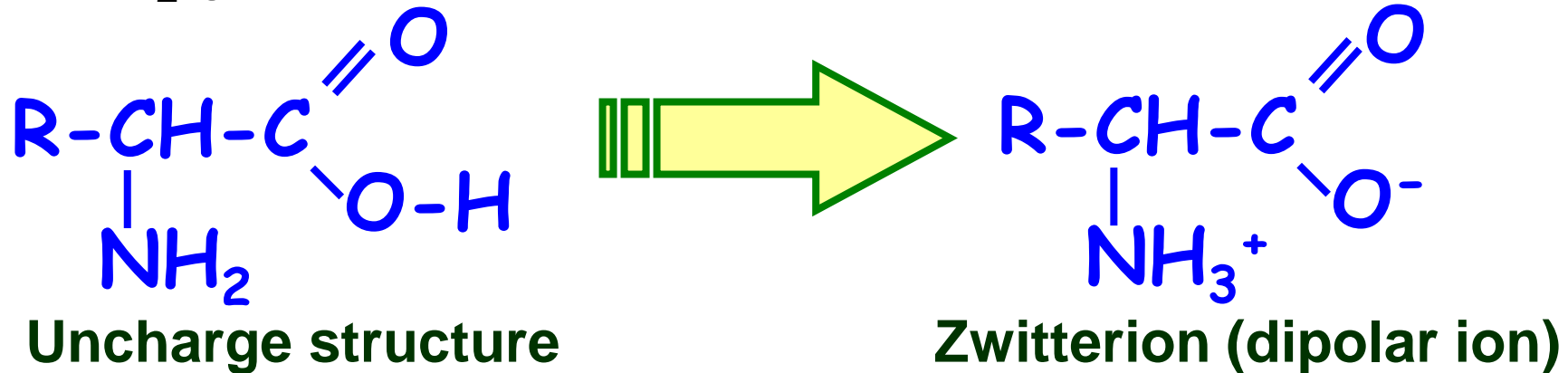
α -Amino acids are dipolar ions. The term used for dipolar ion is zwitterion .

Remarks!!

Dipolar ions = zwitterion

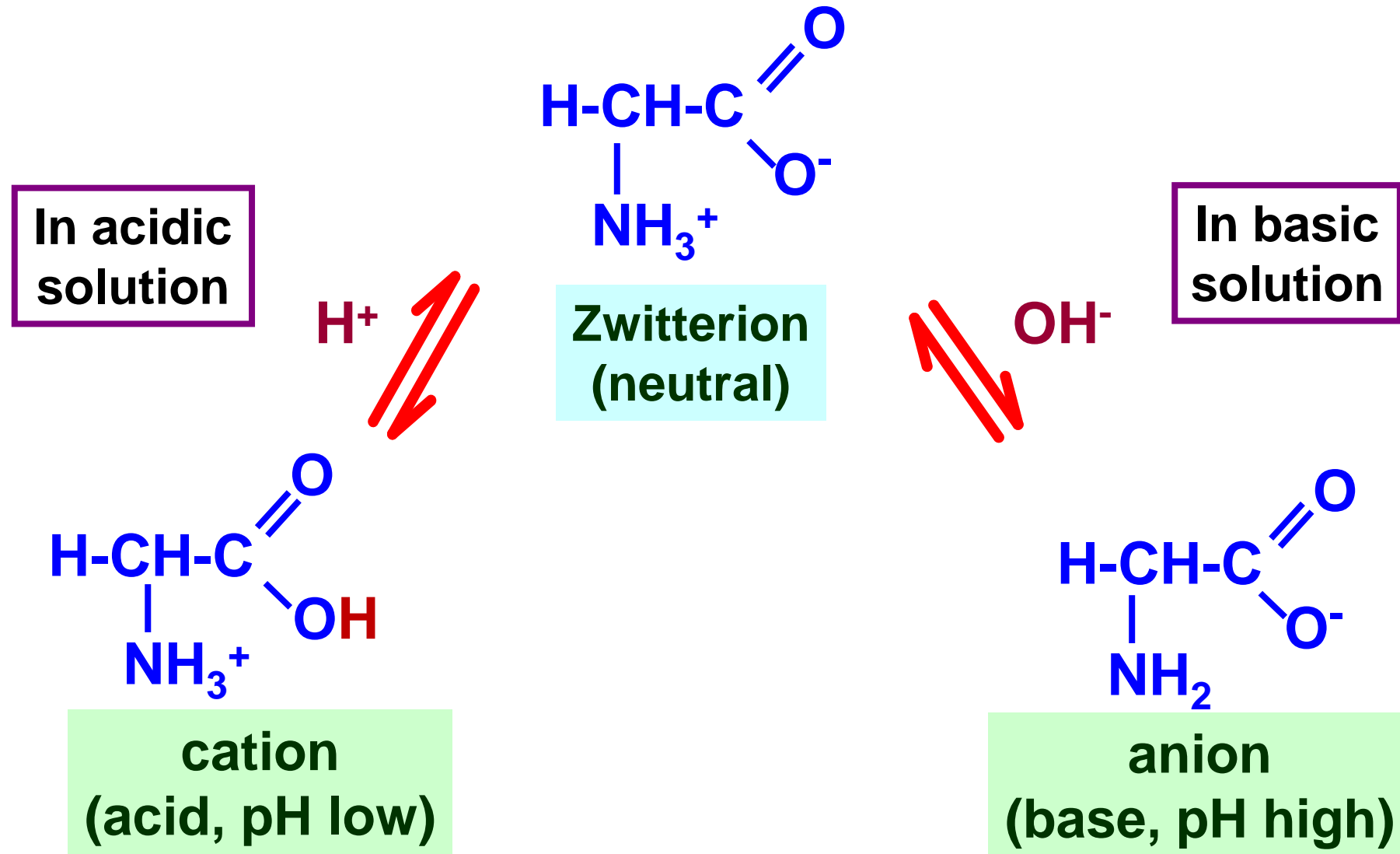
Zwitterion

- ❑ Zwitterion is a molecule with an overall charge of zero which has a **positive charge at one part** and a **negative charge group** at another.
- ❑ In **neutral solution** and **in solid state**, amino acids exist as zwitterion.
- ❑ A zwitterion formed when a proton from the -COOH group is donated to the -NH_2 group of the same molecule.



However, the molecules has no net charge

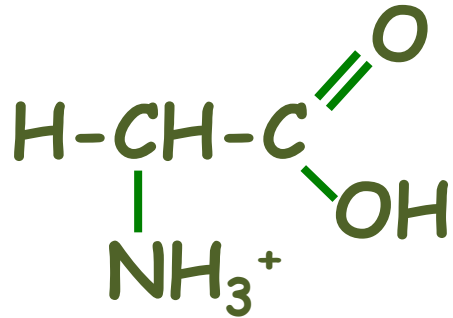
- @ Neutral amino acids are amphoteric.
- @ It can react as acid or base depending on the pH of the solution.



Amino acids can exist in 3 forms depending on the pH of the solution.

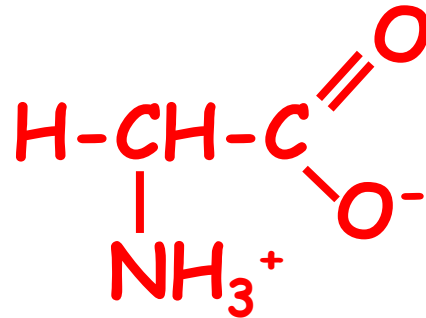
In ACIDIC

Cations are
predominates



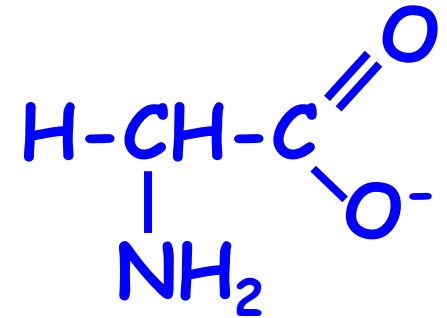
In NEUTRAL

Zwitterions are
predominates



In BASIC

Anions are
predominates



Isoelectric point, pI

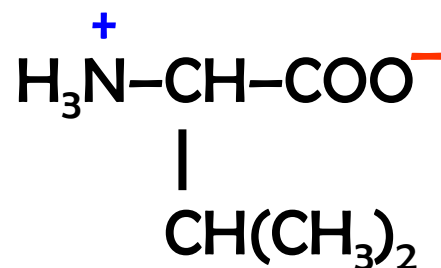
- ❑ The pH whereby the concentration of zwitterion is at maximum and there is equal concentration of both anion and cation.
- ❑ At pI, zwitterion has no net charge.

- Each amino acids has its own specific pI.
- For example: Alanine
 - ✓ Isoelectric point for Alanine is at pH 6.02
 - ✓ At this pH, Alanine exist as zwitterion

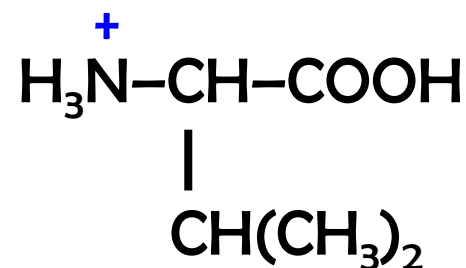
Example :

The isoelectric point of valine, 2-amino-3-methylbutanoic acid is 6.0. Predict the structural formula of valine at

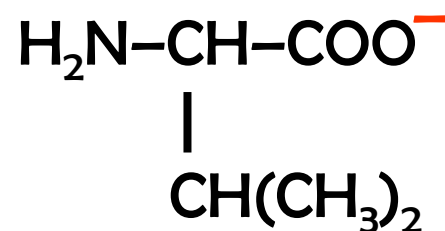
a) Isoelectric point, pI (pH 6)



c) pH 1



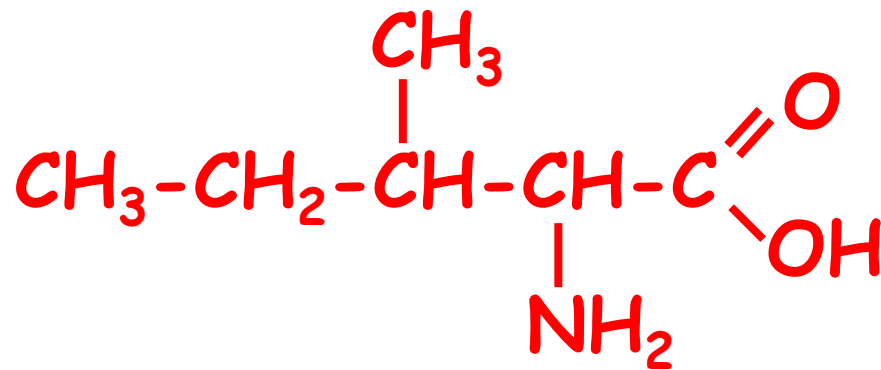
b) pH 12



PAST YEARS QUESTIONS:

Isoleucine, 2-amino-3-methylpentanoic acid is a α -amino acid. Its isoelectric point is 6.0. Draw the structural formula of isoleucine at pH 1.0, pH 6.0 and pH 10.0.

PROBLEM SOLVING:



2-amino-3-methylpentanoic
acid

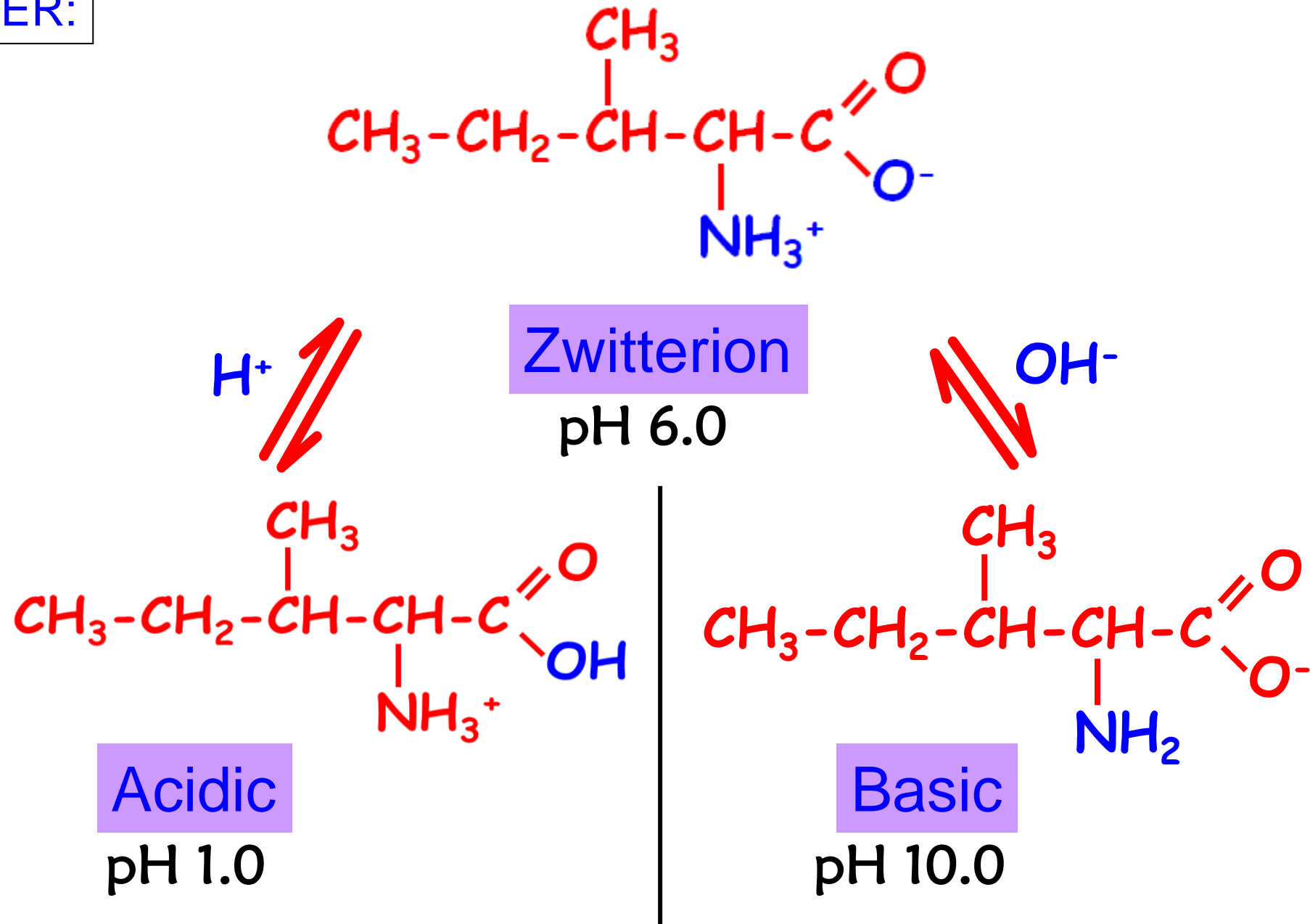
pH 6.0 – Isoelectric point
– zwitterion (neutral)

pH 1.0 – acidic solution

pH 10.0 – basic solution



ANSWER:



12.2 : CHEMICAL PROPERTIES

Students should be able to:

- a) Explain the reaction of amino acid with:
 - i. hydrochloric acid
 - ii. sodium hydroxide
 - iii. nitrous acid
 - iv. alcohols in the presence of an acid catalyst. (C3)
- b) Explain the formation of peptide bond in dipeptides. (C3)

12.2: CHEMICAL PROPERTIES

Since amino acid have both **acidic ($-\text{COOH}$)** and **basic ($-\text{NH}_2$)** functional group, amino acids will undergo 2 main types of chemical reactions:

1. Reactions characteristic of the amino group

- ✓ Reaction with HCl
- ✓ Reaction with nitrous acid (HNO_2)

2. Reactions characteristic of the carboxyl group

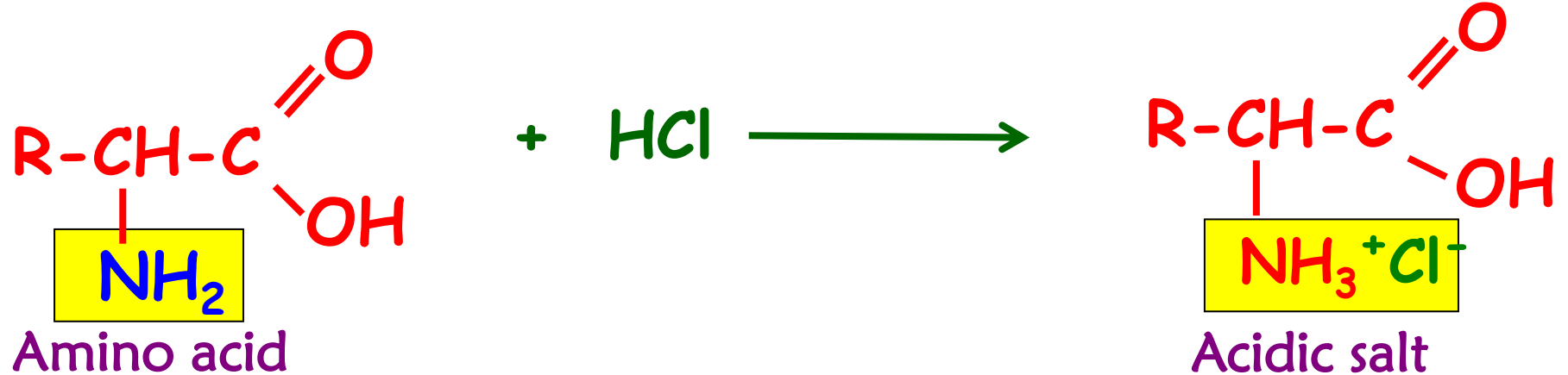
- ✓ Reaction with NaOH
- ✓ Reaction with alcohols in the presence of an acid catalyst (esterification)

Reaction of amino group (-NH₂)

① Reaction with HCl

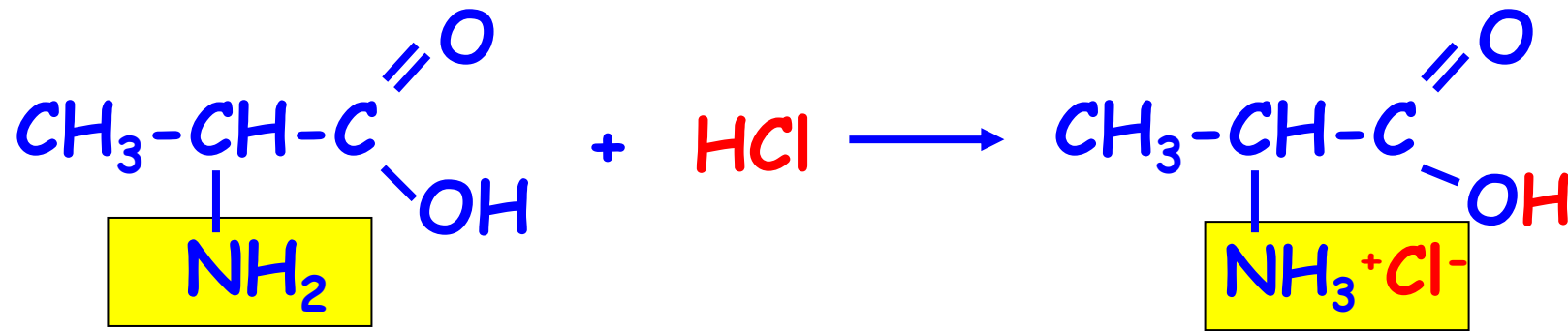
+ Amino group is basic, so it reacts with HCl to form salt.

+ General equation:



Example:

Reaction of Alanine with HCl



Alanine

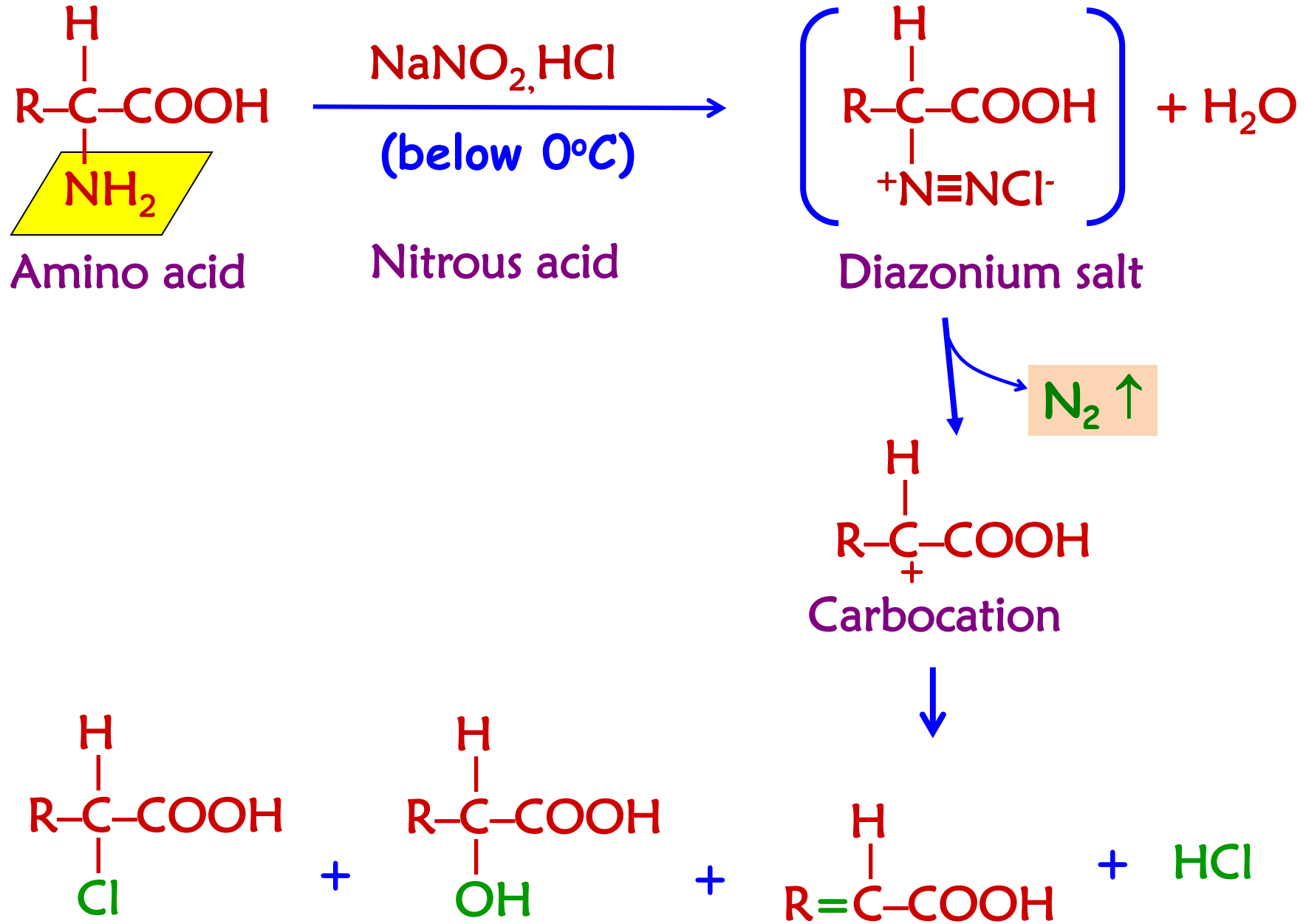
② Reaction with Nitrous Acid (HNO_2)

- ✚ Nitrous acid are prepared *in situ* from $\text{NaNO}_2 + \text{HCl}$
- ✚ Amino group react with HNO_2 at 0°C to form diazonium salt as intermediate product
- ✚ This salt decompose easily by losing N_2 to form carbocation.
- ✚ The carbocation react with nucleophile to form a mixture of products such as **alcohol, alkene, haloalkanes** and HCl

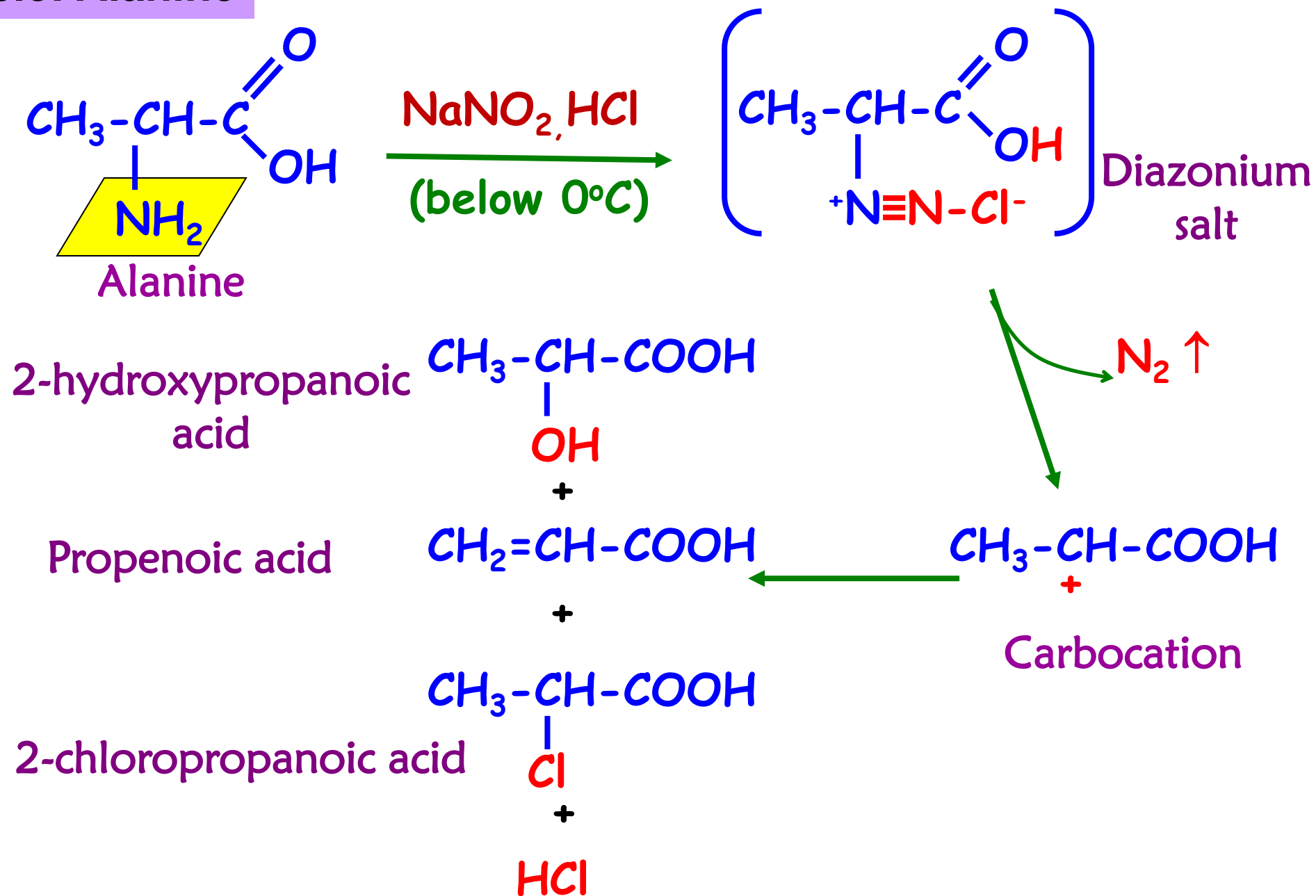
Same reaction that we
learn in 1° aliphatic
amine with nitrous acid



General Equation:



+ Example: Alanine

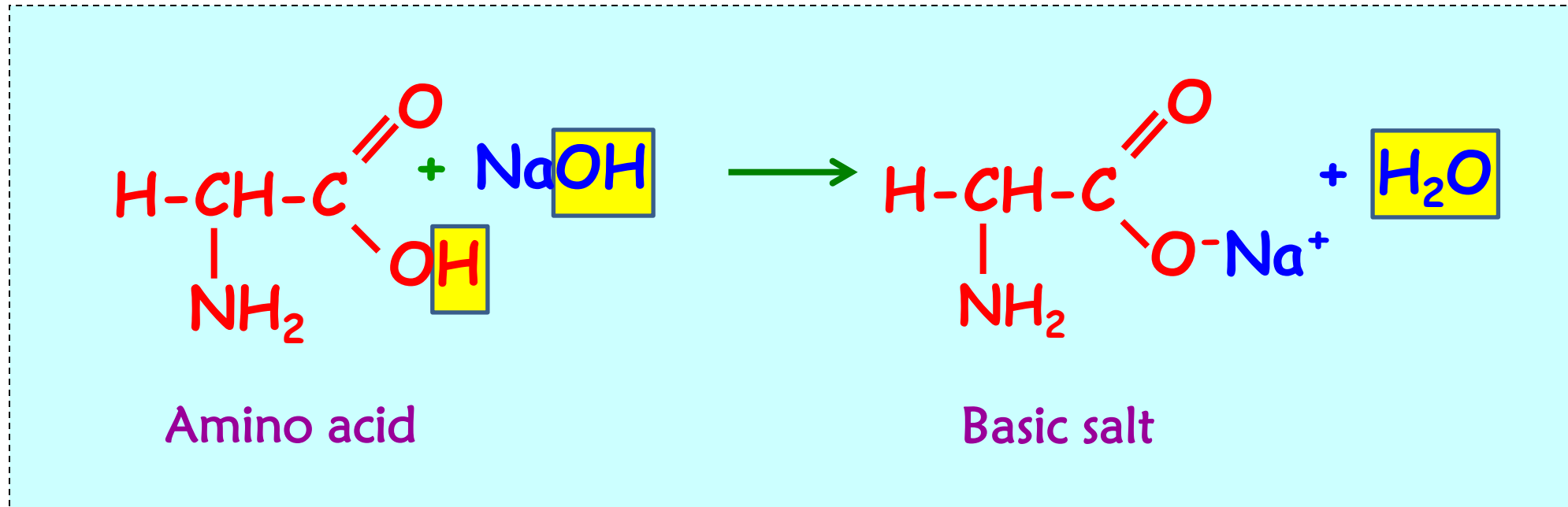


Reaction of Carboxyl group (-COOH)

① Reaction with Sodium Hydroxide (NaOH)

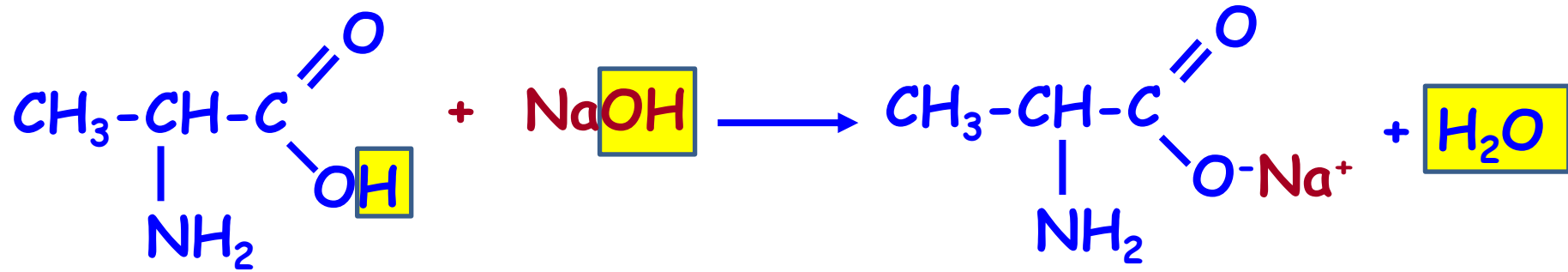
+ Carboxyl group is acidic, so it reacts with NaOH to form salt.

+ **General equation:**



Example:

Reaction of Alanine with NaOH



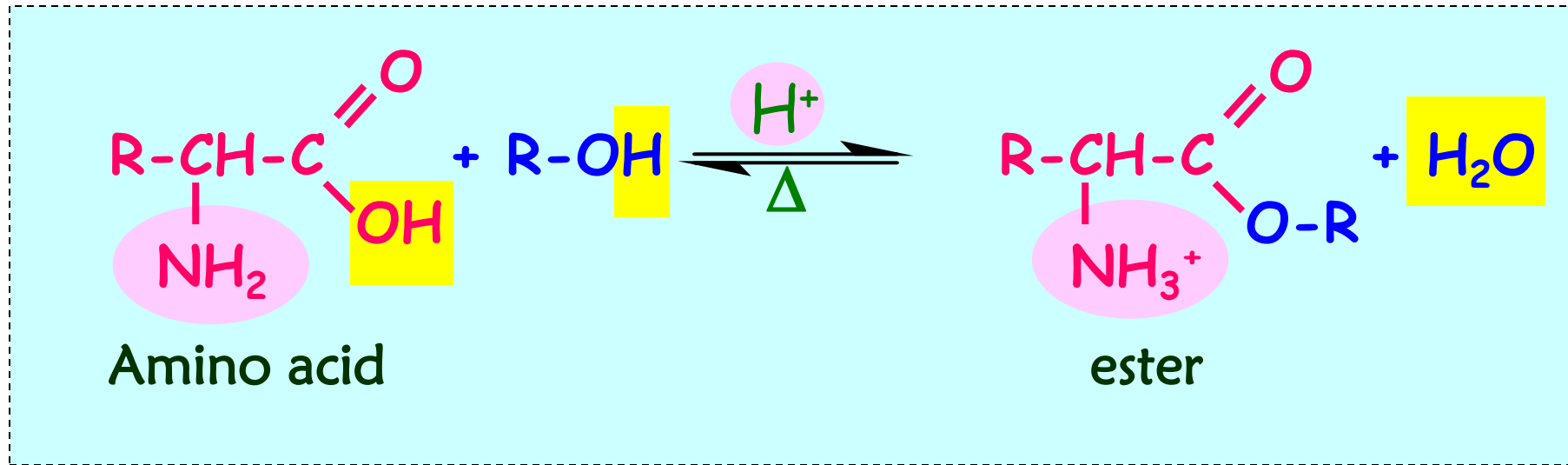
2-aminopropanoic acid

sodium 2-aminopropanoic acid

② Reaction with alcohol (esterification)

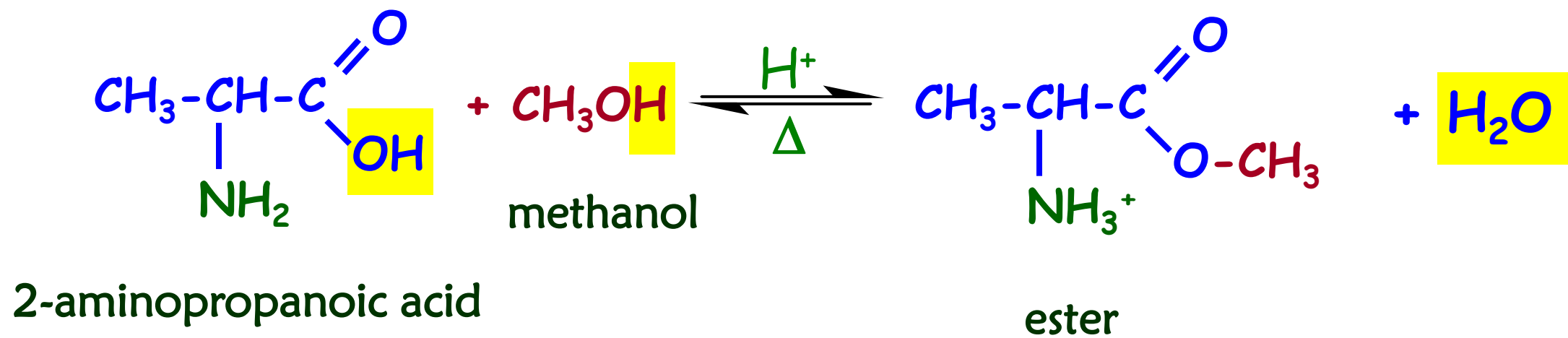
- Amino acid reacts with alcohol in presence of an acid (normally HCl/H₂SO₄) as a catalyst to form ester and water

General equation:



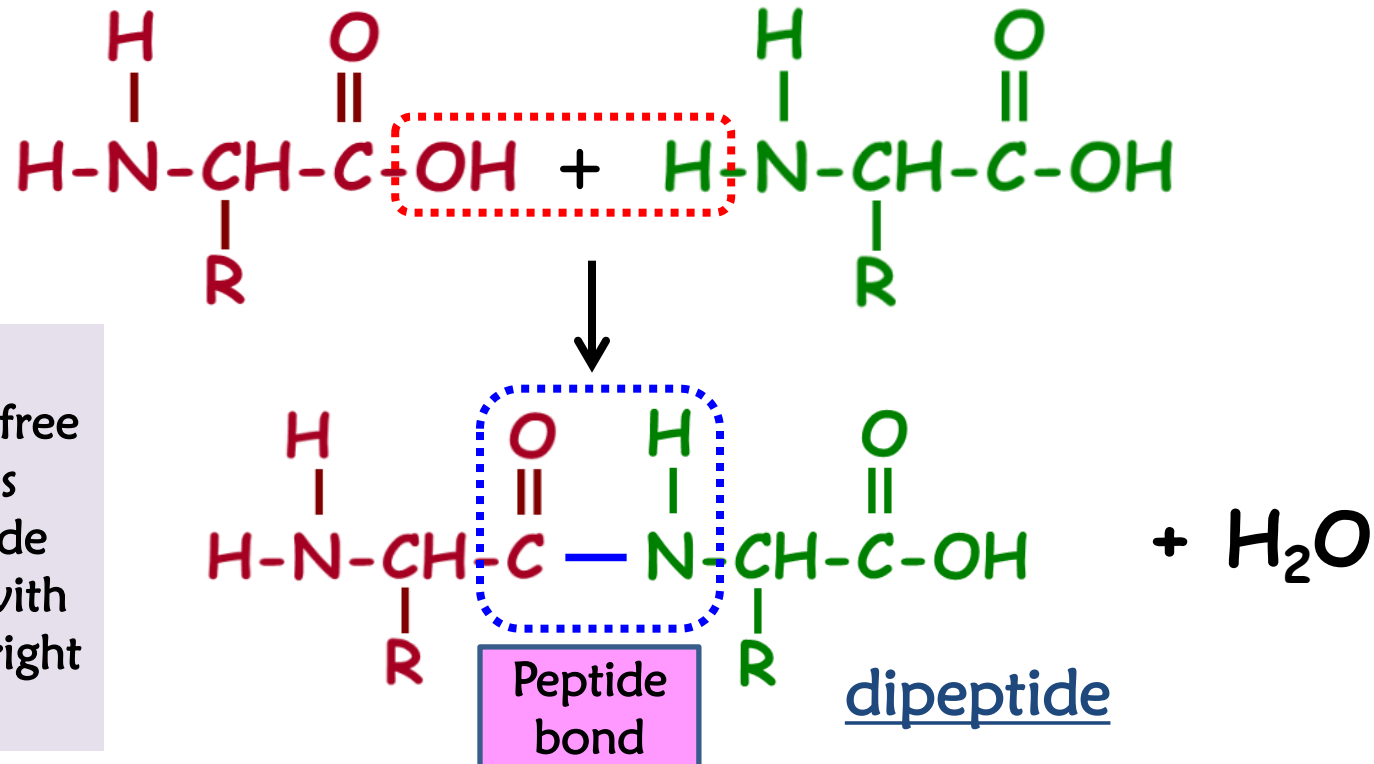
Example:

Alanine reacts with methanol in presence of HCl



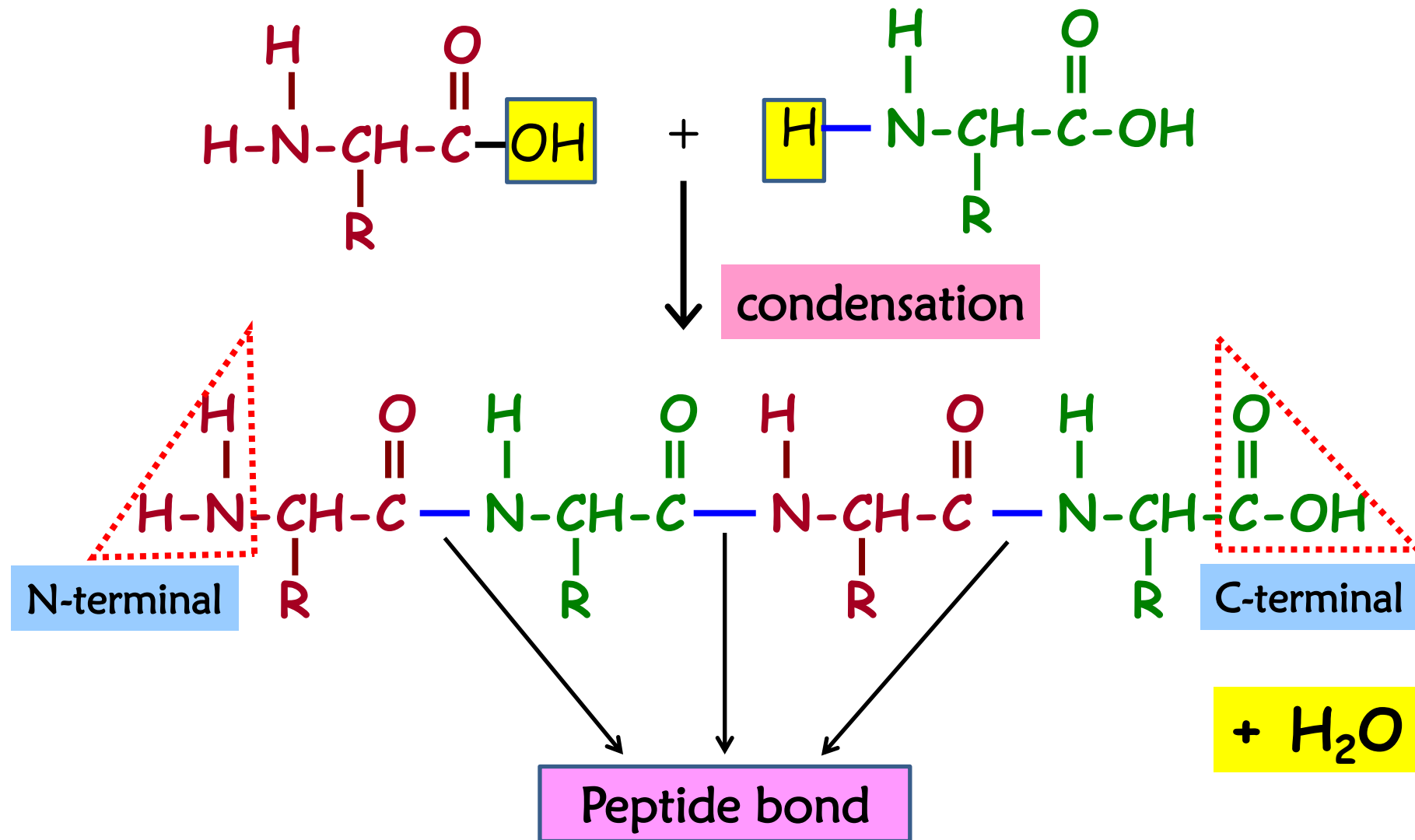
Formation of peptide bonds

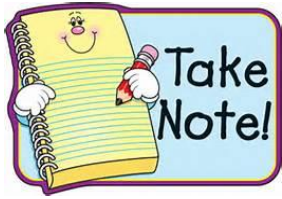
- ✚ **Peptide bond** is a linkage between two amino acid residue.
- ✚ When 2 amino acids react together, H_2O is eliminated, and a compound formed is known as dipeptide



NOTES:

N- terminal (with free $-\text{NH}_2$) is always written on left side and C-terminal (with free $-\text{COOH}$) on right side





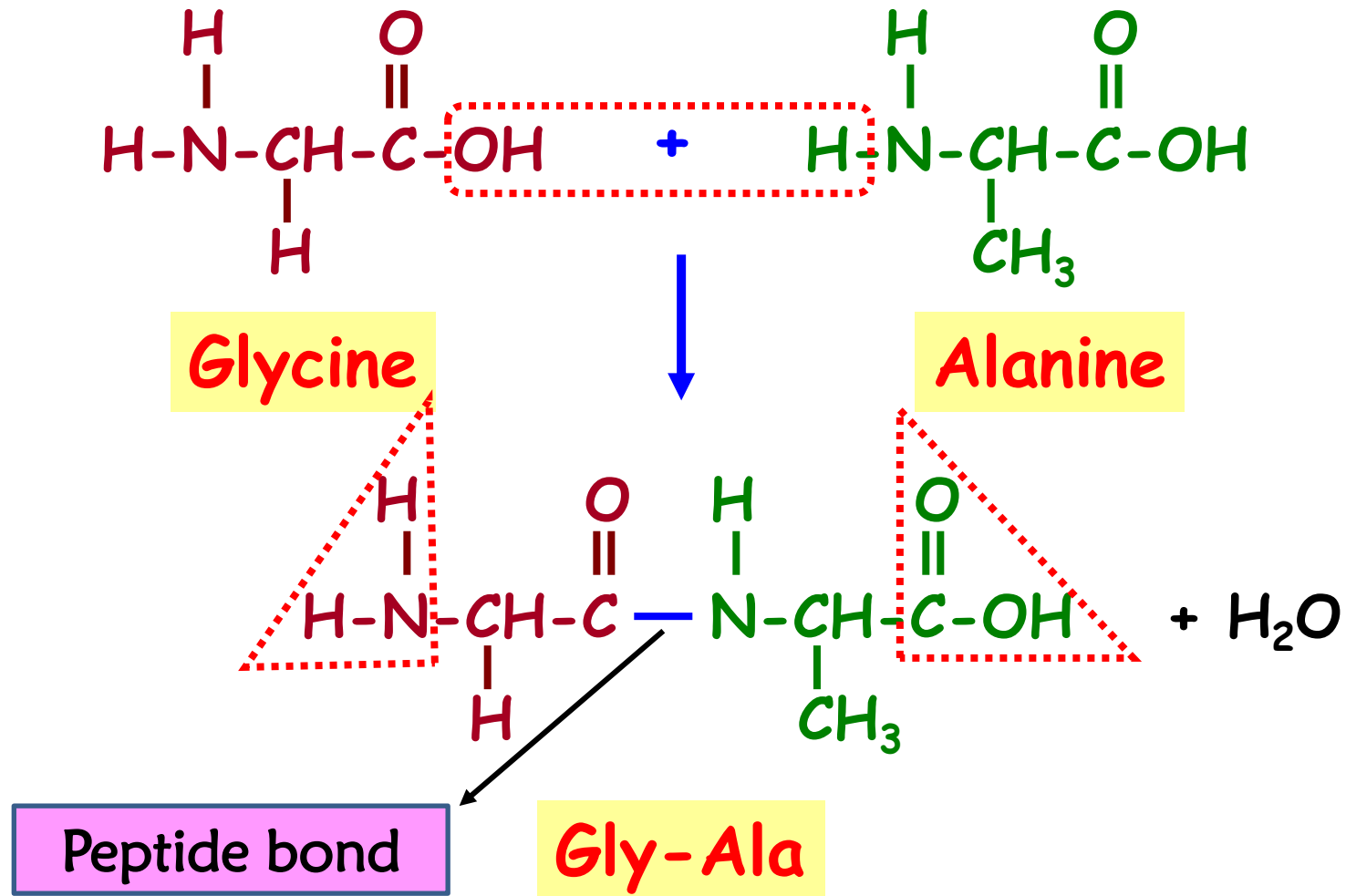
+ 2 amino acids = dipeptide

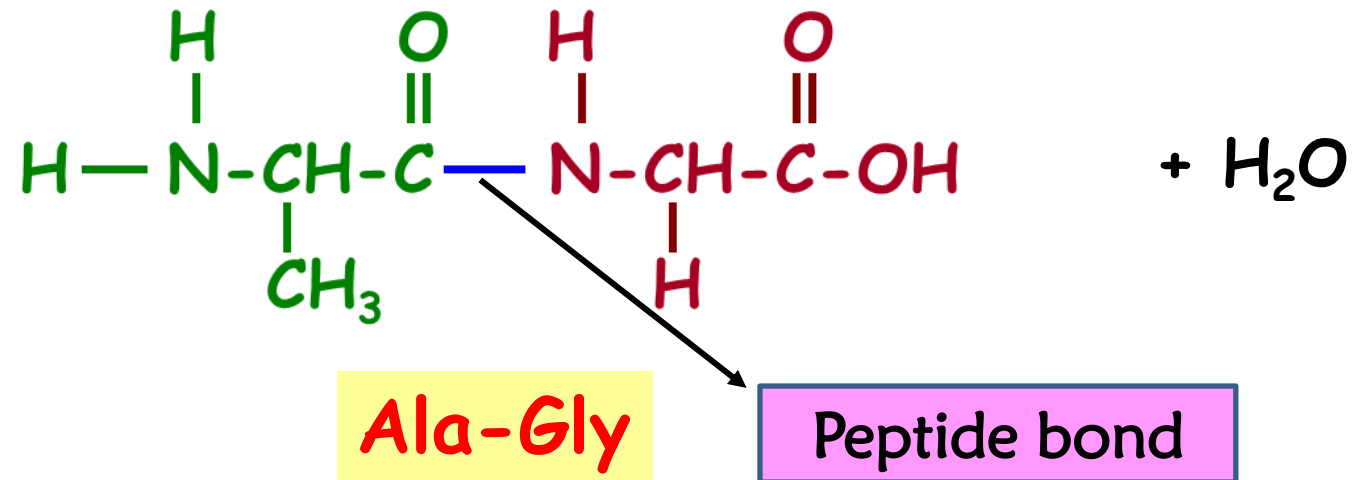
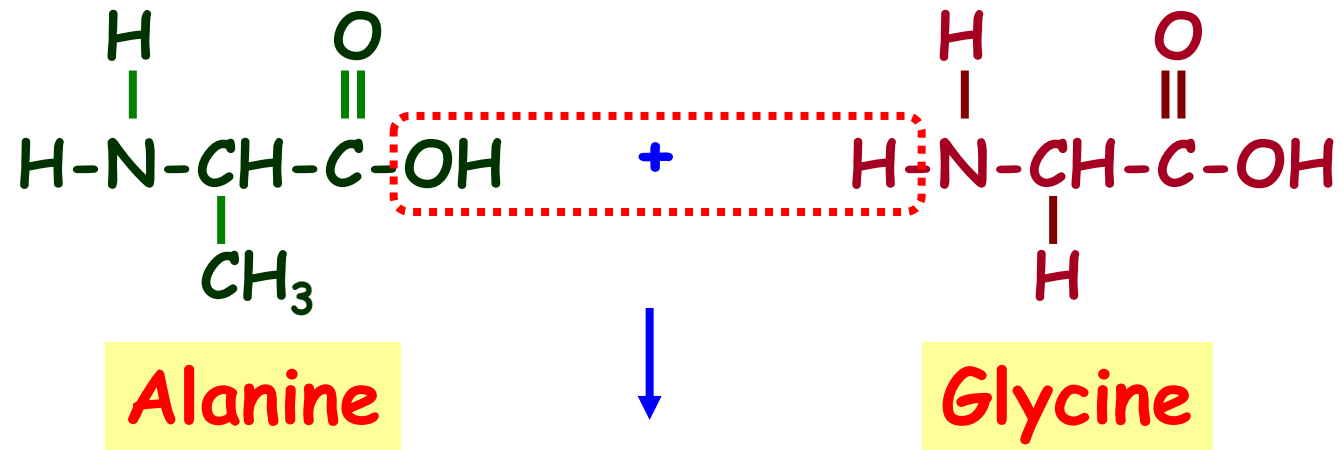
+ 3 amino acids = tripeptide

+ 15-30 amino acids = oligopeptide

+ > 30 amino acids = polypeptide

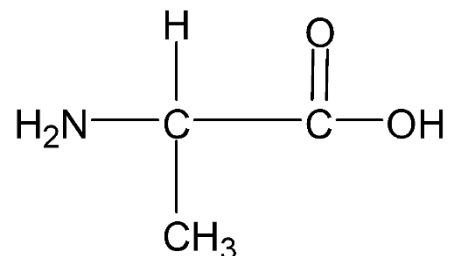
- ✚ Example: Formation of two dipeptides from Glycine (Gly) and Alanine (Ala)
- ✚ The name of the dipeptides are glycylalanine (Gly-Ala) and alanylglycine (Ala-Gly)





Exercise:

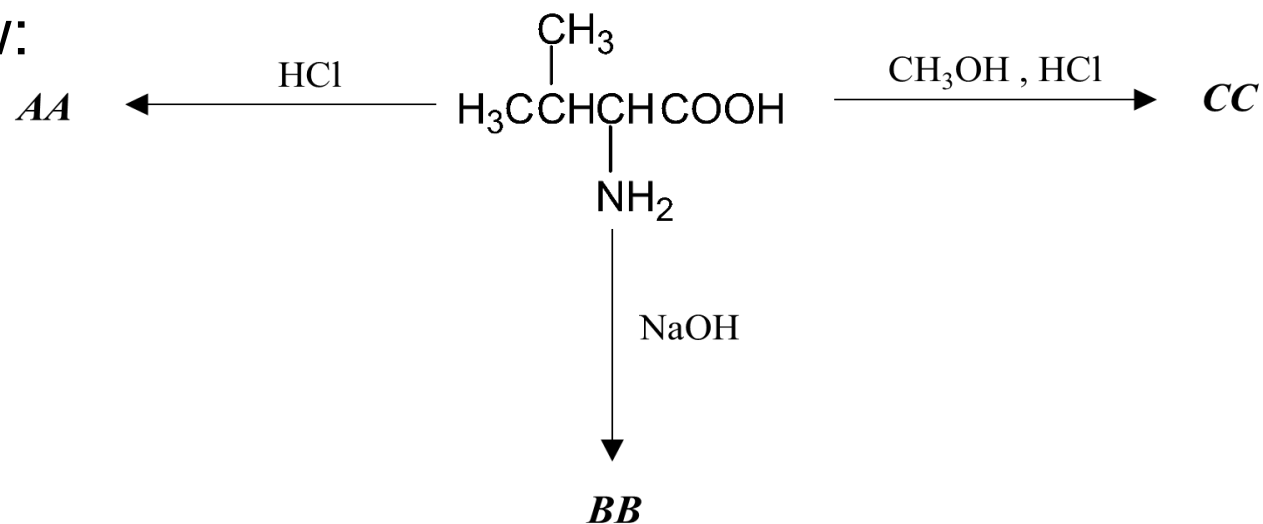
(a) The structure of Alanine is given as below:



Draw the structure of Alanine as zwitterion.

[1 mark]

(b) Reaction of valine with several reagents are shown below:



Draw the structures of products **AA**, **BB** and **CC**.

[3 marks]