

Protein metabolism

SYNOPSIS

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4.METABOLISM OF AMMONIA

5.UTILIZATION OF AMMONIA

6.TRANSPORT OF AMMONIA

7.UREA CYCLE

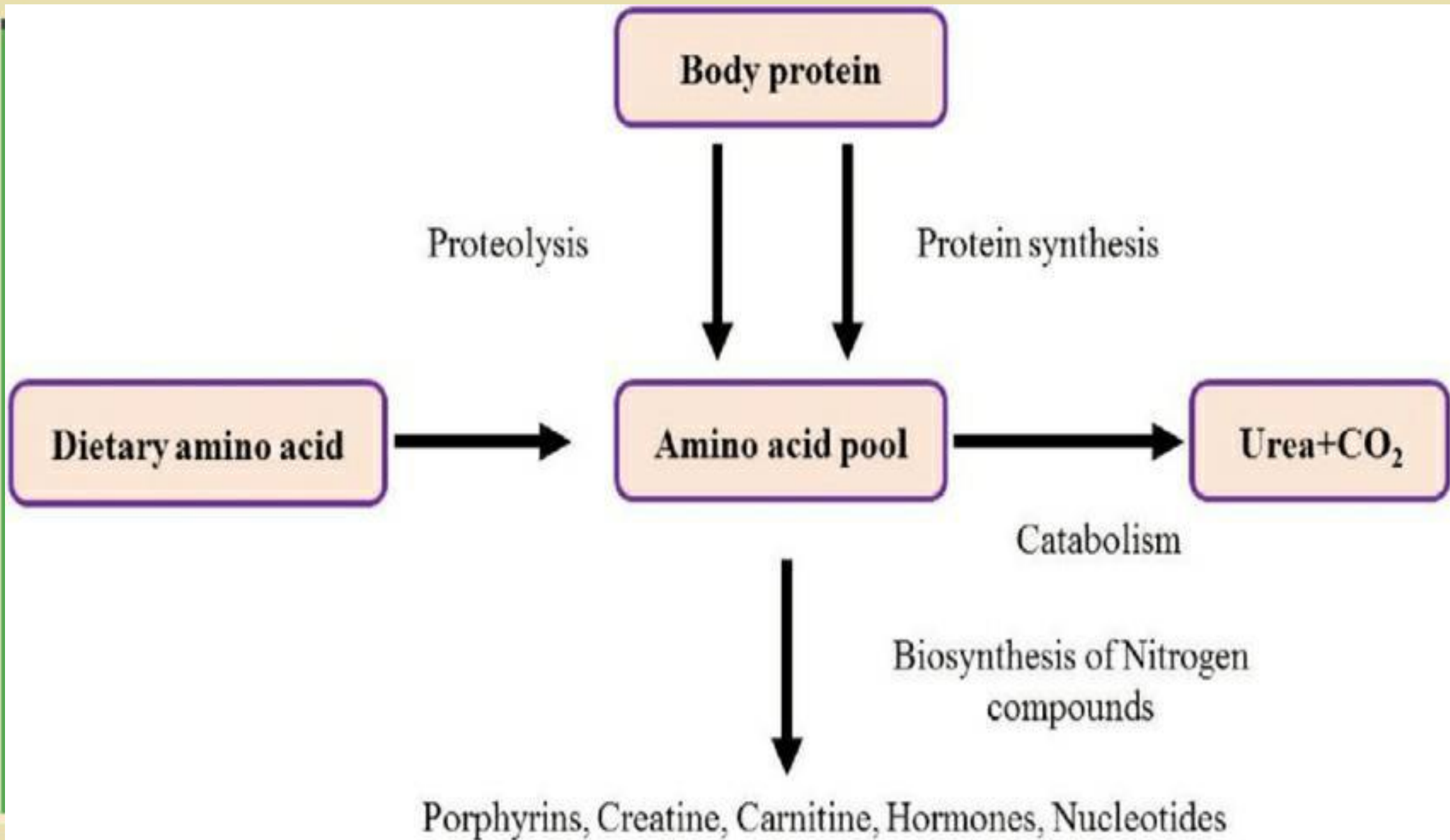
8.REGULATION OF UREA CYCLE.

Introduction – protein metabolism Denotes the various biochemical Processes Responsible for the Synthesis of protein and amino acids the breakdown of proteins by catabolism.

Protein turnover

In cell biology, **protein turnover** refers to the replacement of older proteins as they are broken down within the cell.

Different types of proteins have very different turnover rates



Amino acid pool

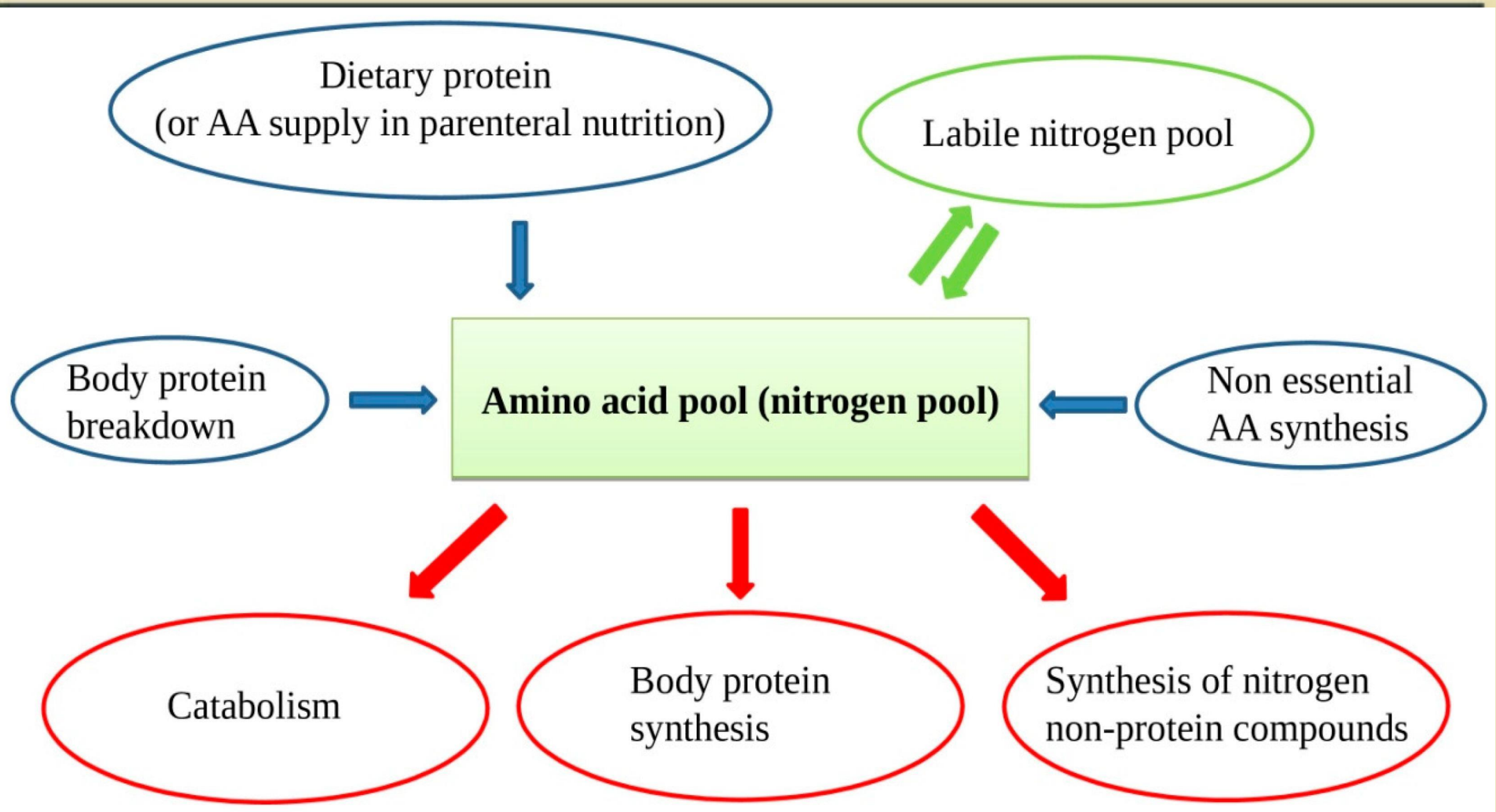
- Amino acid released by dietary and tissue protein
- Mix with free amino acids of body constitutes = 100 gm.
- Glutamate , glutamine – 50%
- Essential amino acids – 10
- Remaining Non – Essential amino acids

- Proteins rich in proline , Glutamate ,serine and Threonine are rapidly degraded and have short half lives

Amino acid pool -: Amino acid released by

1. Dietary and
2. Tissue Protein .

- There is no storage form of amino acid like glycogen and Triglyceride .
- Excess intake of proteins (Amino acid) are metabolised then oxidised to provide energy or converted glucose or fat.
- Amino groups lost as Urea – Excreted.



Metabolism of Ammonia

- **SOURCES OF AMMONIA**

1. Amino acid synthesis protein ,
2. Protein degraded to amino acid,
3. From liver -: (a) Transamination
(b) Oxidative deamination
4. From kidney -: Glutaminase reaction
5. From Intestine -: By bacterial Action
6. From Diet -: Aminos

7. From Catabolism -: Purines (Adnine)

Pyrimidine (Cytosine)

8. From - non Oxidative : Deamination : Aminoacids

UTILIZATION OF AMMONIA

- Glutamate + Ammonia. \rightarrow Glutamine
- Glutamine synthetase : Liver Brain and Kidney
- Brain-: Major Mechanism for removal of Ammonia is Glutamate formation
- $\alpha\text{KG} + \text{NH}_3 + \text{NADPH} + \text{H}^+ \rightarrow \text{Glutamate} + \text{NADP}^+$
- Glutamate may be considered as a major transport form of NH_3 from tissue to liver .

TRANSPORT OF AMMONIA

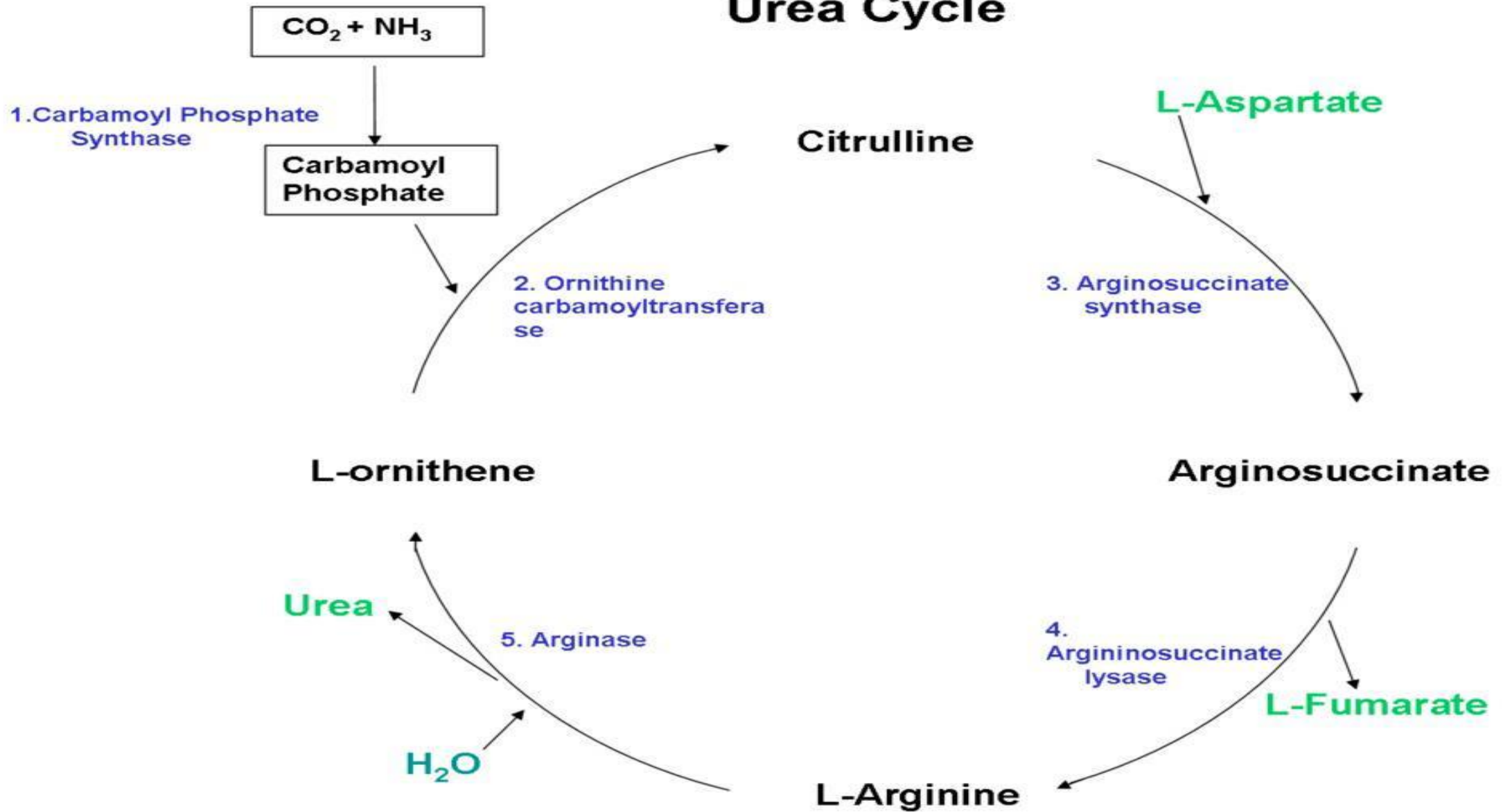
- Ammonia is constantly Produced in tissue
- Plasma Ammonia – 10-20 $\mu\text{g}/\text{dl}$
- Elevated levels cause symptoms of Ammonia Intoxication
- SYMPTOMS -: Tremor, Slurring of speech
Blurring of vision \rightarrow Coma and death.

UREA CYCLE

- Enzymes of urea cycle
- Regulation of urea cycle
- Energetics of urea cycle
- Clinical significance of blood urea
- Disorders of urea cycle

- Urea cycle is also called as krebs Henseleits or Omithine cycle.
- Site -: Liver.
- Urea synthesized in liver released into blood cleared by kidneys.
- Urea cycle delivered into five steps

Urea Cycle



Nitrogen atoms of urea are delivered from ammonia and alpha amino group of aspartic acid .

One mole of urea synthesis requires 4 mole of ATP

Step 1. Glutamate ,glutamine- 50%.



Carbomoyl phosphate synthase 1 (CPS-1)

It is mitochondria enzymes .

Allosteric Activater is **N- Acetyl Glutamate** .

Step 2. Formation of Citrulline.

- Carbomoyal Phosphate + Ornithine



Ornithine Transcarbomoylase.

Citrulline+ pi

- Ornithine trans carbomoylase is also a mitochondria enzymes.

This step onwards the reactions occurred in cytoplasm.

- Step 3. Formation of Argininosuccinate.
- Citrulline + Aspartate + ATP .



Argininosuccinate Synthase

Argininosuccinate + AMP + ppi

Step 4. Formation of Arginine .

Argininosuccinate



Argininosuccinase

Arginine + Fumarate .

Step 5. Formation Of Urea

Arginine + H₂O \longrightarrow Ornithine + Urea .

Regulation Of Urea Cycle .

(A) Carbamoyl Phosphate Synthase – 1

Allosteric activator – N – Acetylglutamate more Glutamate , more Acetylglutamate , more CPS-1 Activity, leads more Urea synthesis .

(B) During Starvation , urea cycle enzyme activities are increased to meet the demands of increased rate of protein Catabolism.

Thank You

