Lactic Acidosis

Dr. Sumbul Fatma

1 Lecture

Cardiovascular Block

Overview

- Introduction to metabolic acid-base disorders
  - Metabolic acidosis and alkalosis
- Lactic acidosis
  - Definition
  - Lactate metabolism in tissue
  - Mechanisms involved in lactic acidosis
  - Types and causes of lactic acidosis
  - Diagnosis and treatment

Metabolic acid-base disorders

Changes in bicarbonate conc. in the extracellular fluid (ECF) cause acid-base disorders

- Occur due to high conc. or loss of H^+ ions
- Can lead to:
  - Metabolic acidosis
  - Metabolic alkalosis

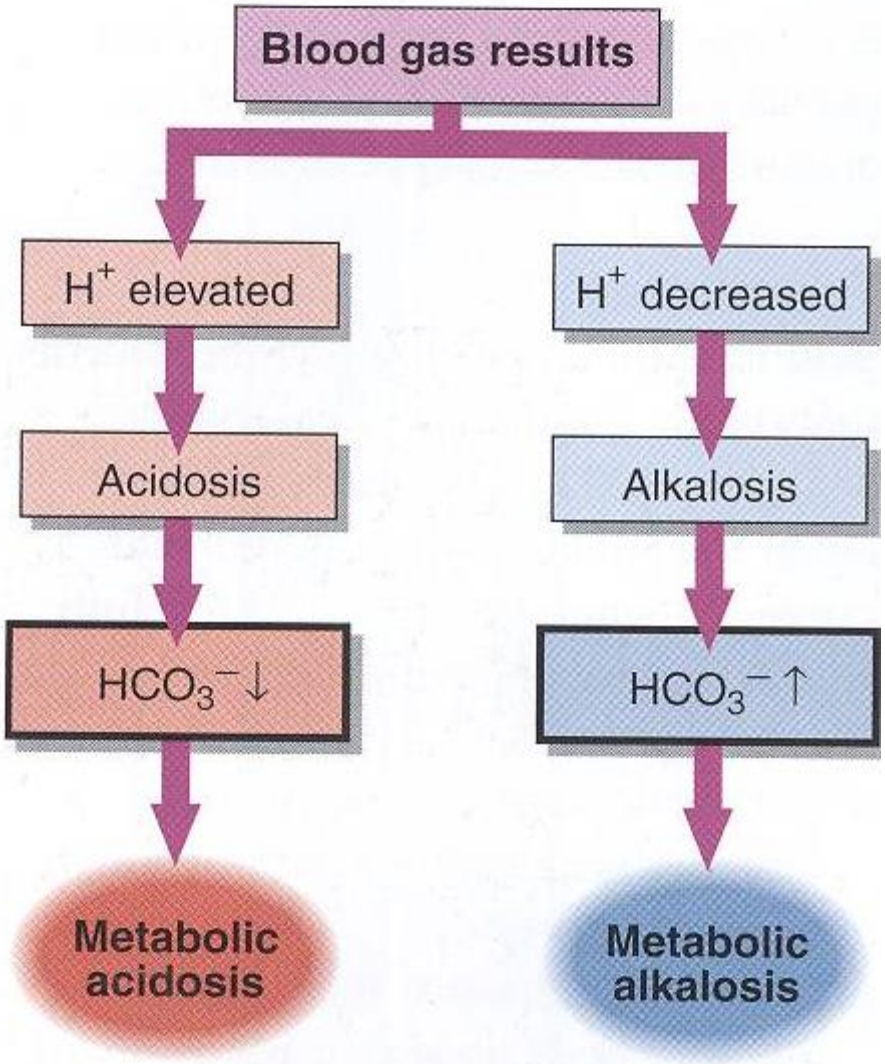


Fig. 1 **Recognizing primary metabolic acid–base disorders by inspecting the HCO_3^- concentration.**

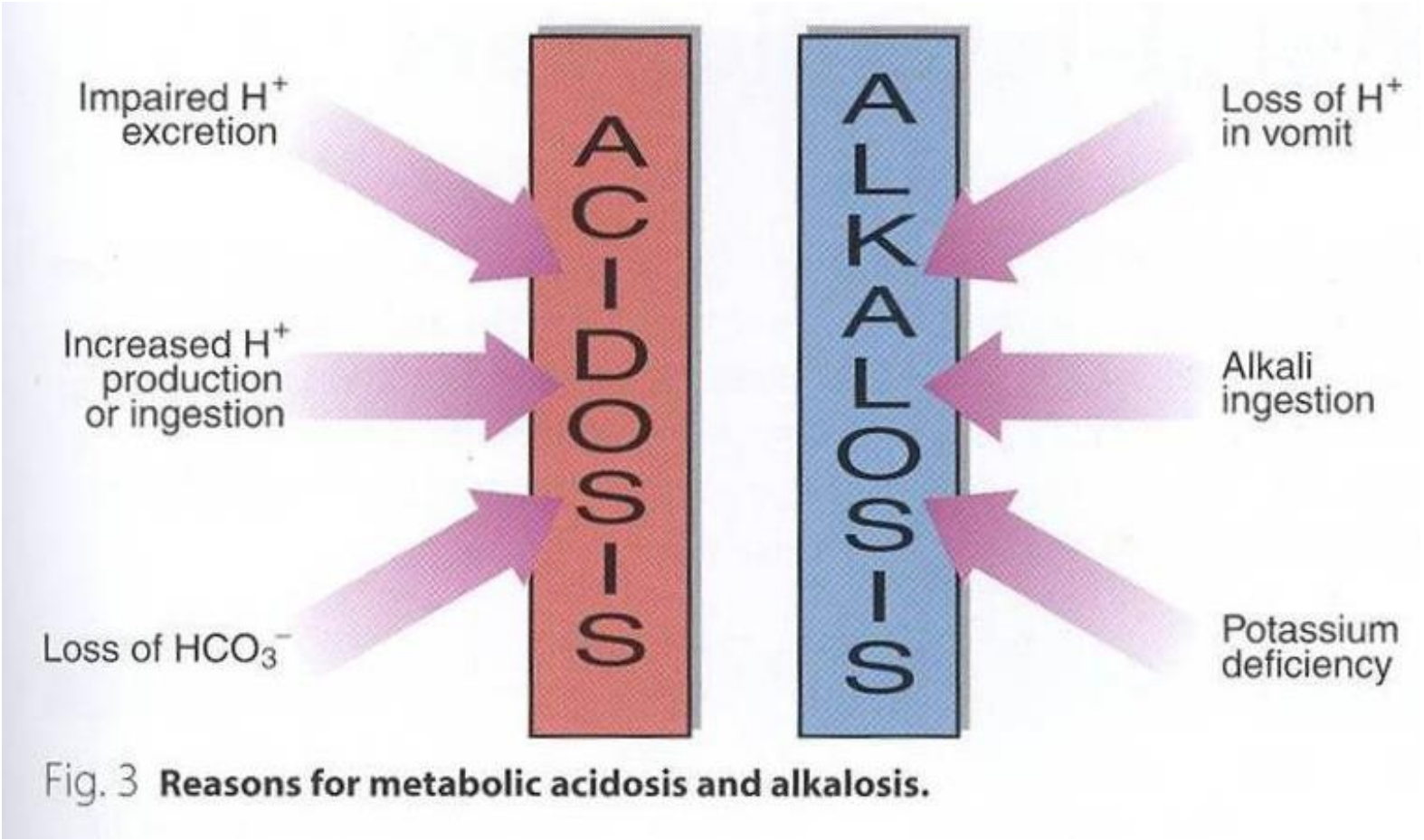


Fig. 3 **Reasons for metabolic acidosis and alkalosis.**

Metabolic acidosis

Reduction in bicarbonate conc. of ECF

Causes are:

- Increased production of H^+ ions
- Ingestion of H^+ or drugs metabolized to acids
- Impaired excretion of H^+

Anion gap

- It is the difference between the sum of:
  - Na^+ and K^+ (cations) and
  - the sum of Cl^- and HCO_3^- (anions)
- Helps in assessing acid-base problems
- Normal anion gap: 3-11 mEq/L
- High anion gap: >11 mEq/L (acidosis)
- Low anion gap: <3 mEq/L (alkalosis)

Metabolic acidosis

High anion gap occurs in:

- Renal disease
- Diabetic ketoacidosis
- Lactic acidosis
- Chronic diarrhea
- Poisoning
- Renal tubular acidosis

Clinical effects of acidosis

Hyperventilation is the compensatory physiological response to acidosis

- Increased H^+ conc. stimulates respiratory response
- Hyperventilation: deep, rapid, and gasping respiratory pattern
- Arrhythmia, cardiac arrest
- Loss of consciousness, coma, death

Metabolic acidosis

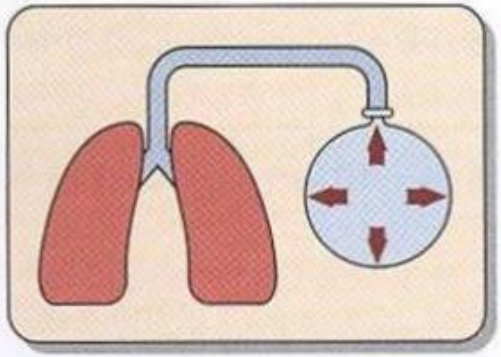
$$[H^+]\uparrow \propto \frac{PCO_2}{[HCO_3^-]\downarrow}$$

Acidosis develops

$$[H^+]\uparrow \propto \frac{PCO_2\downarrow}{[HCO_3^-]\downarrow}$$

Respiratory compensation occurs quickly

Increased ventilation



Metabolic alkalosis

Increase in bicarbonate conc. in ECF

Causes are:

- Loss of H^+ ions in gastric fluid due to vomiting
- Ingestion of sodium bicarbonate
- Potassium deficiency as a result of diuretic therapy

Clinical effects of alkalosis

- Hypoventilation (depressed breathing)
  - Increases PCO_2 to compensate alkalosis
  - Respiratory arrest
- Confusion, coma, death

Metabolic alkalosis

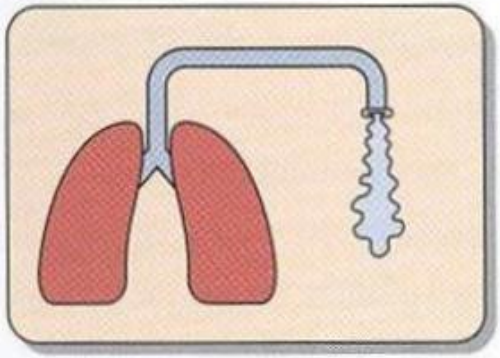
$$[H^+]\downarrow \propto \frac{PCO_2}{[HCO_3^-]\uparrow}$$

Alkalosis develops

$$[H^+]\downarrow \propto \frac{PCO_2\uparrow}{[HCO_3^-]\uparrow}$$

Respiratory compensation occurs quickly

Decreased ventilation



Lactic acidosis

- Elevated conc. of plasma lactate is called lactic acidosis
- Occurs either due to:
  - Failure of circulatory system (hypoxia)
  - Disorders of carbohydrate metabolism

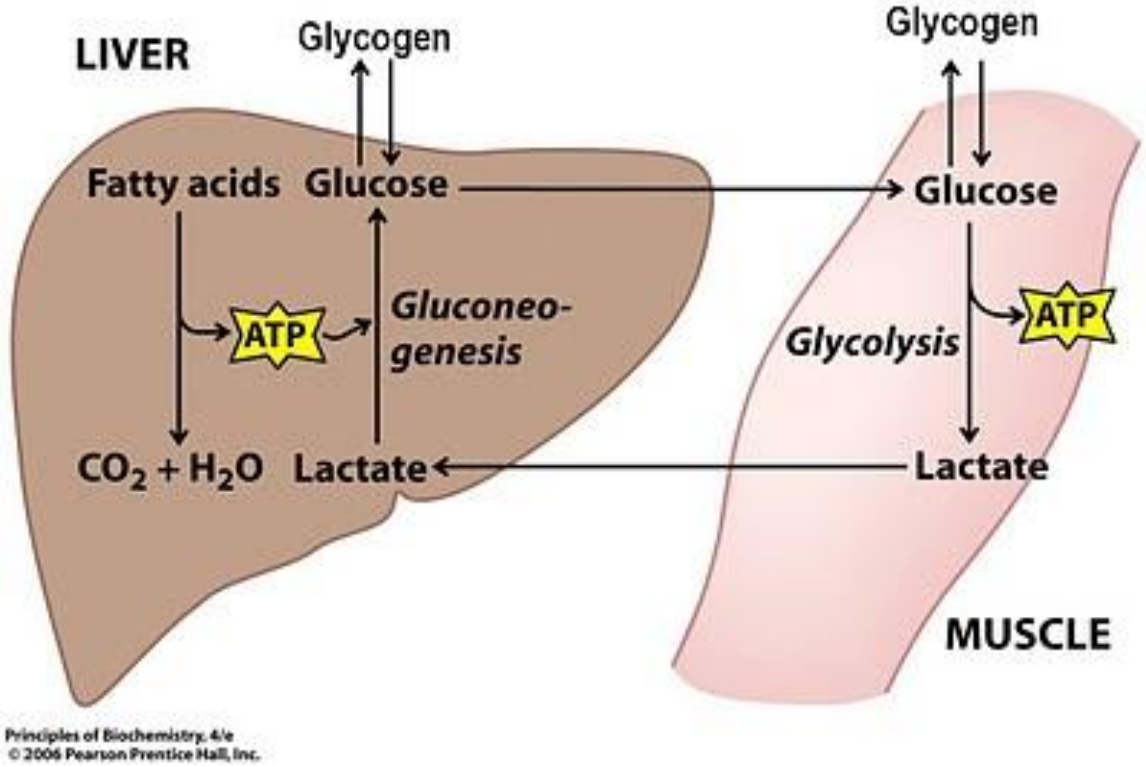
Lactate metabolism in tissue

- The body tissues produce ~ 1500 mmoles of lactate each day
- The lactate enters blood stream and metabolized mainly by the liver (Cori cycle)
- All tissues can produce lactate under anaerobic conditions
- Pyruvate is converted to lactate by lactate dehydrogenase enzyme

Pyruvate + NADH + H^+

Lactate dehydrogenase

Lactate + NAD^+



The Cori cycle

Lactate metabolism in tissue

- The skeletal muscles produce high amounts of lactate during vigorous exercise
- Lactate is metabolized in liver (60%) and kidney (30%) to glucose
- Some lactate is metabolized to CO2 and water (Krebs cycle)

Mechanisms involved in lactic acidosis

Lactic acidosis can occur due to:

- Excessive tissue lactate production
- Impaired hepatic metabolism of lactate

Types and causes of lactic acidosis

Type A

- Due to hypoxia in tissues (most common)
- Hypoxia causes impaired oxidative phosphorylation and decreased ATP synthesis
- To survive, the cells switch to anaerobic glycolysis for ATP synthesis
- This produces lactate as a final product
- The amount of oxygen required to recover from oxygen deficiency is called oxygen debt

Types and causes of lactic acidosis

- Type A is due to inadequate supply of oxygen to tissues in:
  - Myocardial infarction
  - Pulmonary embolism
  - Uncontrolled hemorrhage
  - Tissue hypoperfusion (shock, cardiac arrest, acute heart failure, etc.)
  - Anaerobic muscular exercise

Types and causes of lactic acidosis

Type B

- Due to disorders in carbohydrate metabolism
  - Congenital lactic acidosis is due to deficiency of pyruvate dehydrogenase enzyme
- Chronic hepatic disease accompanied by shock or bleeding
- Liver failure
- Drug intoxication

Diagnosis and treatment

- Diagnosis done by measuring blood lactate levels
  - Hyperlactemia: 2 – 5 mmols/L
  - Severe lactic acidosis: > 5 mmols/L
- Treatment:
  - Correcting the underlying conditions
  - Restoring adequate tissue oxygen
  - Avoiding sodium bicarbonate