

Acid-base balance disorders – practical interpretation

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Learning outcomes

- 🕒 Student will learn to detect the presence of acid-base balance disorder.
- 🕒 Student will learn how to evaluate the compensation of acid-base balance disorder.
- 🕒 Student will learn how to detect unmeasured anions of organic acids and to suggest differential diagnosis of the cause of metabolic acidosis.

Content of the lecture

- Basic terms and definitions
- Classification of acid-base balance disorders and it's evaluation
- Evaluation of compensation of acid-base balance disorders,
Boston rules
- Calculation of anion gap
- Differential diagnosis of metabolic acidosis
- Practical examples

Basic terms and definitions

- Acid-base balance – the set of processes leading to the formation and disappearance of acids and bases in the internal environment
- Constant concentration of hydrogen cations H^+ (40 nmol/l)
- (norm 7,35 – 7,45)
- **Acid** – a substance that is capable of releasing H^+ → pH reduction
- **Base** – a substance that is capable of accepting H^+ → pH increasement

Basic terms and definitions

- ⦿ **Acidemia** – increase in H^+ concentration – pH drop below 7,35
- ⦿ **Acidosis** – processes in the internal environment that lead to the formation of acids or to the disappearance of bases and thus to a decrease in pH
- ⦿ **Alkalemia** – decrease in H^+ concentration – pH increase above 7,45
- ⦿ **Alkalosis** – processes that lead to the disappearance of acids or to the formation of bases and thus to a rise in pH

Basic terms and definitions

• Buffer - a solution containing a conjugated pair of a weak acid and its salt, maintaining a stable pH



Henderson-Hasselbalch equation:

Basic terms and definitions



pK of carbonic acid

CO₂ solubility coefficient

The change in pH is determined by the partial pressure of CO₂ (PaCO₂) and the concentration of bicarbonate (HCO₃⁻).

Classification of ABB disorders

Respiratory

- Caused by changes in PaCO_2 (norm PaCO_2 4,6-6 kPa)
- Acidosis – insufficient CO_2 elimination, pCO_2 in extracellular fluid rises
- Alkalosis – alveolar ventilation higher than needed to maintain normal PaCO_2 , pCO_2 decreases

Metabolic

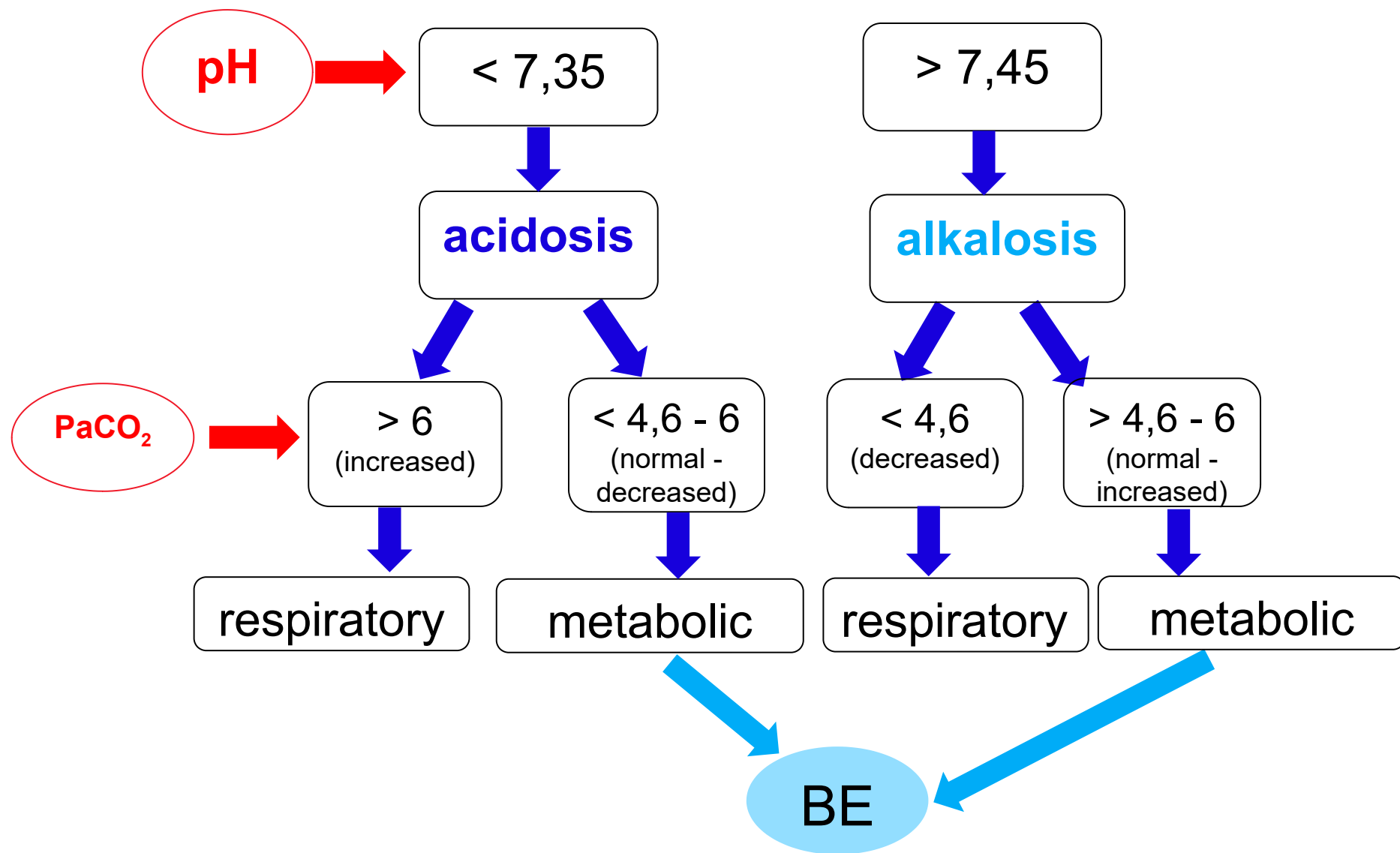
- Caused by changes in HCO_3^- (norm 24 ± 2 mmol/l), changes in the concentration of buffer bases (BB)

Excess of bases – Buffer Base (BB)

- ☉ Sum of base concentrations of all buffering systems
- ☉ Bicarbonate bases, non-bicarbonate bases (phosphates, albumin)

Base Excess (BE)

- ☉ The difference between the current BB value and the normal BB value
- ☉ The amount of acid or base that must be added to the plasma to achieve a normal pH of 7,4 under a constant of PaCO_2 5 kPa
- ☉ Norm -2 – +2
- ☉ **Metabolic acidosis**: decrease in BE < -2 or decrease in HCO_3^- concentration
- ☉ **Metabolic alkalosis**: increase in BE > 2 or increase in HCO_3^- concentration



Compensation of acid-base balance disorders

- If the $\text{PaCO}_2/\text{HCO}_3^-$ ratio is kept constant, the pH will be normal
- Primary ABB disturbance → reaction by opposing change to normalize pH → **compensation = secondary ABB disturbance**
- Respiratory system – PaCO_2 regulation
- Metabolic regulation - kidneys, liver

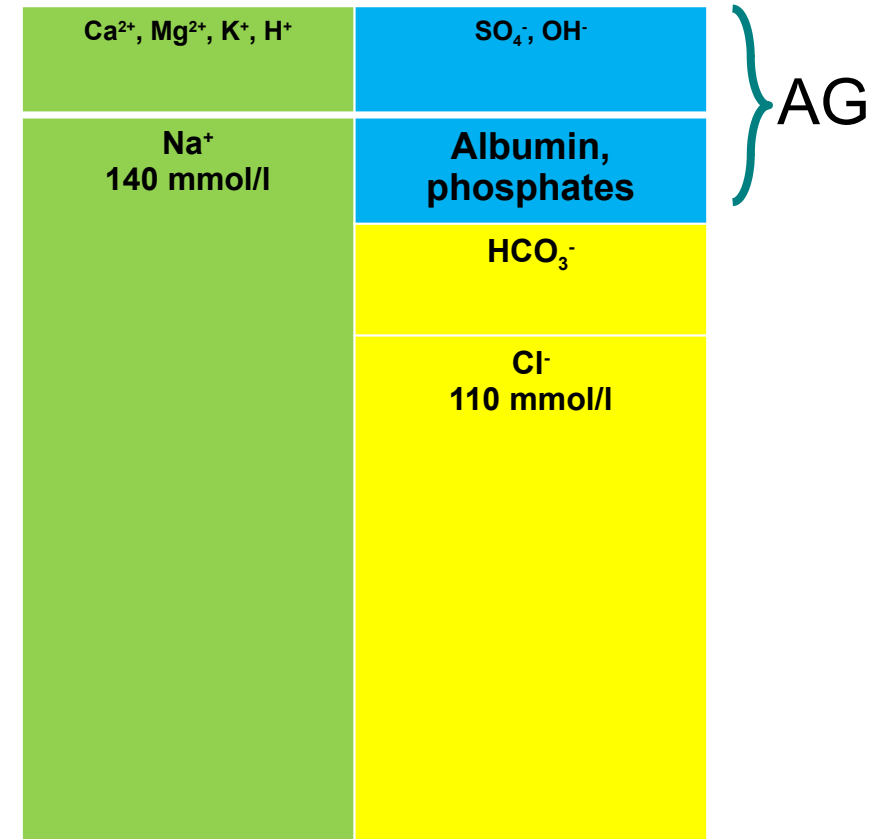
! Normal pH does not mean that an ABB disorder is not present
! ABB disturbance is present if there is an abnormal pH or PaCO_2

Degree of ABB disturbance compensation – Boston rules

Primary disorder	HCO ₃ ⁻ concentration (mmol/l)	PaCO ₂ (kPa)
Metabolic acidosis	< 22	HCO ₃ ⁻ /5 + 1
Metabolic alkalosis	> 26	HCO ₃ ⁻ /10 + 2,6
Acute respiratory acidosis	(PaCO ₂ – 5,3) x + 24 increased by per kPa	> 6
Chronic respiratory acidosis	(PaCO ₂ – 5,3) x 3 + 24 increased by 3 per kPa	> 6
Acute respiratory alkalosis	24 – 1,5 x (5,3 – PaCO ₂) decreased by 1,5 per kPa	< 4,7
Chronic respiratory alkalosis	24 – 4 x (5,3 – PaCO ₂) decreased by 4 per kPa	< 4,7

Anion Gap (AG)

- Detection of the presence of acid anions undetectable by conventional laboratory methods
- Law of conservation of electroneutrality - the sums of all positive and negative charges in the system are equal
- **AG = (Na⁺ + K⁺) – (Cl⁻ + HCO₃⁻)**
- norm 12 4 mmol/l



Corrected anion gap – AGc

Ca ₂ ⁺ , Mg ₂ ⁺ , K ⁺ , H ⁺	SO ₄ ⁻ , OH ⁻	} AG
	Albumin, phosphates	
Na ⁺ 140 mmol/l	A ⁻	} AGc
	HCO ₃ ⁻	
	Cl ⁻ 110 mmol/l	

- Takes into account the current concentration of albumin and phosphate
- $AGc = [(Na^+ + K^+) - (Cl^- + HCO_3^-)] - (0,2 \times \text{albumin (g/l)} + 1,5[\text{phosphate (mmol/l)}]) - \text{lactate}$
- Norm AGc = 0

→ MAC with a high AG (HAGMA)
→ MAC with a normal AG

Metabolic acidosis

With a high AG - HAGMA

- M Methanol
- U Uremia
- D Diabetic/alcoholic ketoacidosis
- P Propylene glycol
- I Iron, isoniazid
- L Lactate
- E Ethylene glycol/ethanol
- S Salicylates

With a normal AG - hyperchloremic

- 1 GIT secretion
- 2 Renal tubular acidosis
- 3 Iatrogenic

Pactical approach:

1. Identify the primary ABB disorder

- Is pH or PaCO₂ abnormal?
- If both pH and PaCO₂ are abnormal, compare the direction of change
- If the change is in the same direction, the primary ABB disorder is metabolic
- If the change is in the opposite direction, the primary ABB disorder is respiratory
- If pH or PaCO₂ is normal, it is a combined disorder

1. Identify the primary ABB disorder

○ Example 1: patient with pH 7,23 and PaCO₂ 3,0 kPa

○ pH PaCO₂ → **primary metabolic acidosis**

○ Example 2: patient with pH 7,23 and PaCO₂ 7,2 kPa

○ pH PaCO₂ → **primary respiratory acidosis**

○ Example 3: patient with pH 7,37 and PaCO₂ 7,5 kPa

○ Normal pH PaCO₂ → **combined respiratory acidosis with metabolic alkalosis**

Practical approach:

2. evaluate the compensatory response

- Boston rules
- Metabolic acidosis: calculate the expected PaCO_2 ($\text{HCO}_3^-/5 + 1$)
- Metabolic alkalosis: calculate the expected PaCO_2 ($\text{HCO}_3^-/10 + 2,6$)
- Respiratory acidosis: calculate the expected HCO_3^- :
 - acute $(\text{PaCO}_2 - 5,3) \times 1 + 24$
 - chronic $(\text{PaCO}_2 - 5,3) \times 3 + 24$
- Respiratory alkalosis: calculate the expected HCO_3^- :
 - acute $24 - 1,5 \times (5,3 - \text{PaCO}_2)$
 - chronic $24 - 4 \times (5,3 - \text{PaCO}_2)$

2. evaluate the compensatory response: example

Patient with COPD exacerbation, dyspnoic, soporous
pH 7,15, PaCO₂ 16,3 kPa, PaO₂ 10,3 kPa, HCO₃ 42 mmol/l,
BE 7,9 mmol/l, SaO₂ 90%

○ Compare the direction of change:

pH PaCO₂ → primary respiratory acidosis

○ Calculate the expected HCO₃ = (PaCO₂ - 5,3) x 3 + 24

$$\text{HCO}_3 = (16,3 - 5,3) \times 3 + 24 = 57$$

○ Compare the actual and calculated HCO₃

→ Primary respiratory acidosis with insufficient compensatory metabolic alkalosis

3. Detection of unmeasurable anions - AG

Example: a patient with a history of chronic alcohol abuse, admitted to hospital in a coma, Kussmaul breathing.

pH 7,31	Na 144 mmol/l
PaCO ₂ 2,9 kPa	K 4,3 mmol/l
HCO ₃ ⁻ 10,6 mmol/l	Cl 106 mmol/l
BE -13,4	Ca ²⁺ 1,12 mmol/l
Albumin 27,8 g/l	Mg 0,81 mmol/l
Lactate 2,4	P 1 mmol/l

1. What is the primary disorder?

pH PaCO₂
HCO₃⁻ significantly negative



Primary metabolic acidosis

2. What is the compensatory response?

pH 7,31	Na 144 mmol/l
PaCO ₂ 2,9 kPa	K 4,3 mmol/l
HCO ₃ ⁻ 10,6 mmol/l	Cl 106 mmol/l
BE -13,4	Ca ²⁺ 1,12 mmol/l
Albumin 27,8 g/l	Mg 0,81 mmol/l
Lactate 2,4	P 1 mmol/l

- Calculate the expected PaCO₂ in case of full compensation
- $\text{PaCO}_2 = \text{HCO}_3^- / 5 + 1$
- $\text{PaCO}_2 = 10,6 / 5 + 1 = 3,12$
- Compare the actual and calculated PaCO₂

○ **Fully compensated respiratory metabolic acidosis**
(no superimposed disorder present)

pH 7,31	Na 144 mmol/l
PaCO ₂ 2,9 kPa	K 4,3 mmol/l
HCO ₃ ⁻ 10,6 mmol/l	Cl 106 mmol/l
BE -13,4	Ca ²⁺ 1,12 mmol/l
Albumin 27,8 g/l	Mg 0,81 mmol/l
Lactate 2,4	P 1 mmol/l

3. Detection of unmeasured anions with AGc

$$\text{AGc} = [(\text{Na}^+ + \text{K}^+) - (\text{Cl}^- + \text{HCO}_3^-)] - (0,2 \times \text{albumin (g/l)} + 1,5[\text{phosphate (mmol/l)}]) - \text{lactate}$$

$$\text{AGc} = [(140 + 4,3) - (106 + 10,6)] - (0,2 \times 27,8) + 1,5 \times 1 - 2,4$$

$$\text{AGc} = 18,24$$

Metabolic acidosis with a high AG \longrightarrow dif. dg.: MUDPILES

Fully compensated respiratory metabolic acidosis with a high AG

Probable cause: alcoholic ketoacidosis or intoxication by toxic alcohols

Take home message

- ABB disturbance is present when there is abnormal pH or PaCO₂.
- When evaluating ABB disorders, first determine what the primary disorder is, then evaluate the compensatory response.
- BE is a more reliable parameter to diagnose metabolic acidosis than bicarbonate concentration.
- The anion gap (AG) calculation is used to detect unmeasurable anions that cause metabolic acidosis.

Source literature

- Maláska J, Stašek J, Kratochvíl M, Zvoníček V. *Intenzivní medicína v praxi*. Praha: Maxdorf, [2020]. Jessenius. ISBN 978-80-7345-675-7.
- McNamara J, Worthley, LIG. Acid-base balance: part I. Physiology. *Critical care and resuscitation*. 2001;3:181-187.
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