

## TOPIC 5.A. Transport Phenomena

Spatial heterogeneity of values of system parameters (temperature, pressure, concentration, electric potential, etc.) results in transport phenomena. Transfer physical quantity occurs in the opposite direction of the gradient, and the system ultimately will pass to the equilibrium state.

**Exercise 5.1a.** Define the following terms:

*Flux* of some quantity is \_\_\_\_\_

*Flux density* is \_\_\_\_\_

*Gradient* of any physical quantity is \_\_\_\_\_

**Exercise 5.2a.** Fill in the table:

Transport phenomenon	The parameter which determines the transport	Law (name)	Equation
Diffusion			(5.1a)
Viscosity			(5.2a)
Thermal conductivity			(5.3a)

**Exercise 5.3a.** Specify arrows the accordance between the physical quantity and the formula for its calculation:

1) Diffusion coefficient	a) $\eta = \frac{1}{3} \langle \lambda \rangle \langle v \rangle \rho [Pa \cdot s]$ (5.4a)
2) Coefficient of viscosity	b) $\kappa = \frac{1}{3} \langle \lambda \rangle \langle v \rangle \rho C_{vm} [W \cdot m^{-1} \cdot K^{-1}]$ (5.5a)
3) Thermal conductivity coefficient	c) $D = \frac{1}{3} \langle \lambda \rangle \langle v \rangle [m^2 \cdot s^{-1}]$ (5.6a)

**Exercise 5.4a.** After analyzing the Exercise 5.3 write down the relation between transport coefficients:

$$(5.7a)$$

Transport coefficients determined by the mean free path of molecules, the mean distance that a molecule passes between two collisions:

$$\langle \lambda \rangle = \frac{1}{\sqrt{2} \pi \sigma^2 n_V}, \quad [m] \quad (5.8a)$$

where  $\sigma$  is an effective diameter of molecules (the minimum distance that can get close to centers of two molecules);  $n_V$  is the concentration of molecules.

The average time between two successive collisions of molecules  $\tau$  can be found by dividing the mean

free path of molecules by their mean speed:  $\tau = \frac{\langle \lambda \rangle}{\langle v \rangle}$ . (5.9a)

**Example of problem solution.**

Calculate the time during which air with mass  $m = 720$  mg diffuse from soil to the atmosphere through the surface with area  $S = 1$  m<sup>2</sup>, if the air diffusion coefficient is  $D = 0.04$  cm<sup>2</sup>·s<sup>-1</sup> and density gradient

is  $\frac{\Delta\rho}{\Delta x} = -0.50 \times 10^{-6}$  g·cm<sup>-4</sup>.

**Data:**

$S = 1$  m<sup>2</sup>

$m = 720$  mg =  $720 \times 10^{-6}$  kg

$D = 0.04$  cm<sup>2</sup>·s<sup>-1</sup> =  $0.04 \times 10^{-4}$  m<sup>2</sup>·s<sup>-1</sup>

$\frac{\Delta\rho}{\Delta x} = -0.50 \times 10^{-6}$  g·cm<sup>-4</sup> =

=  $-0.05$  kg·m<sup>-4</sup>

$t = ?$

**Solution:**

The mass of gas transported as a result of diffusion, obeys Fick's law:

$$m = -D \frac{\Delta\rho}{\Delta x} S t$$

Express the time from the Fick's law:

$$t = -\frac{m}{D(\Delta\rho / \Delta x)S}$$

Make calculations of diffusion time:

$$t = -\frac{7.2 \times 10^{-4}}{4 \times 10^{-6} \times (-0.05) \times 1} = 3.60 \times 10^3 \text{ s} = 1 \text{ hour.}$$

**Answer:** Diffusion occurred within one hour.

**Problem 5.5a.** At what pressure the mean free path of hydrogen molecules is 2.5 cm if the gas temperature is 67 °C? The effective diameter of a hydrogen molecule is 0.28 nm.

**Data:**

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**Solution:**

**Answer:** \_\_\_\_\_

**Problem 5.6a.** Calculate how many times dynamic viscosity coefficient of carbon dioxide differs from nitrogen one if both gases are at the same temperature and the same pressure. The effective diameters of gas molecules are also identical.

**Data:**

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**Solution:**

*Answer:* \_\_\_\_\_

**Problem 5.7a.** Calculate the thermal conductivity coefficient of nitrogen gas, if under the same conditions the coefficient of dynamic viscosity for this gas is  $10 \mu\text{Pa}\cdot\text{s}$ .

*Data:*

*Solution:*

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*Answer:* \_\_\_\_\_

**Control questions**

1. Show the interrelation between transport coefficients.
2. Gradient of any physical quantity is...
3. What do you know about viscosity of medicines?
4. What methods of measuring viscosity do you know?
5. Surface tension of liquid medications.
6. Surface-active substances.

**Individual assignments**

1. At what temperature mean free path of air molecules is equal to  $9.3 \times 10^{-8} \text{ m}$ ? The effective diameter of air molecules is  $3 \times 10^{-10} \text{ m}$ . The pressure is normal.
2. Calculate the mean number of collisions per unit of time  $\nu$  molecules of carbon dioxide at  $100 \text{ }^\circ\text{C}$ , if the mean free path of molecules  $\langle \lambda \rangle = 0.03 \text{ nm}$ .
3. Find the thermal conductivity coefficient of hydrogen, if it is known that under the same conditions the coefficient of viscosity for it is equal to  $8.6 \mu\text{Pa}\cdot\text{s}$ .
4. Find coefficient of viscosity of nitrogen under the normal conditions, if its diffusion coefficient is equal to  $1.42 \times 10^{-5} \text{ m}^2\cdot\text{s}^{-1}$ .
5. For what time will amount of heat equals to  $2 \text{ kJ}$  pass through the animal's muscles with cross-section area  $1 \text{ dm}^2$  and thickness  $10 \text{ mm}$  at the temperature is  $38 \text{ }^\circ\text{C}$ . Ambient temperature is  $15 \text{ }^\circ\text{C}$ . Thermal conductivity coefficient of the muscle is  $5.7 \times 10^{-2} \text{ W}\cdot\text{m}^{-1}\cdot\text{K}^{-1}$ .
6. What mass of carbon dioxide will diffuse from soil to atmosphere per 1 hour through the surface of the beds with width  $50 \text{ cm}$  and a length  $18 \text{ m}$  if the visible surface of the beds is 1.5 times less than soil surface obtained by loosening it? Do you need some additional data to solve this problem?
7. For what time will  $720 \text{ mg}$  of carbon dioxide diffuse from black soil into the atmosphere through  $1 \text{ m}^2$  of its surface if density gradient is  $10 \text{ g}\cdot\text{cm}^{-1}$ ? The diffusion coefficient is equal to  $0.04 \text{ cm}\cdot\text{s}^{-1}$ .

## TOPIC 5.B. Transport of Substances Through Biological Membranes

The topic covers the information on the structure of biological membranes, transport of substances through them, generation and propagation of nerve impulse, reception processes and transformations of energy. Biological membranes serve many functions in an organism. They enclose dividing a cell into separate regions, compartments, that allows the cell to maintain a nonequilibrium state of a biosystem. The topic is especially important for pharmacy because the efficiency of medicines greatly depends on their ability to penetrate into tissues that in its turn is determined by permeability of biological membranes to the medicines.

**Exercise 5.1b.** Give detailed answers to the following questions.

Biological membranes have the following properties:

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Semipermeability is \_\_\_\_\_

Membrane fluidity \_\_\_\_\_

Types of membrane lipid motion:

1) Simple motion

- a. \_\_\_\_\_
- b. \_\_\_\_\_
- c. \_\_\_\_\_

2) Composite motion

- a. Lateral diffusion is \_\_\_\_\_  
\_\_\_\_\_
- b. Flip-flop gap is \_\_\_\_\_  
\_\_\_\_\_

In this regard, biological membranes serve the following functions in biosystems:

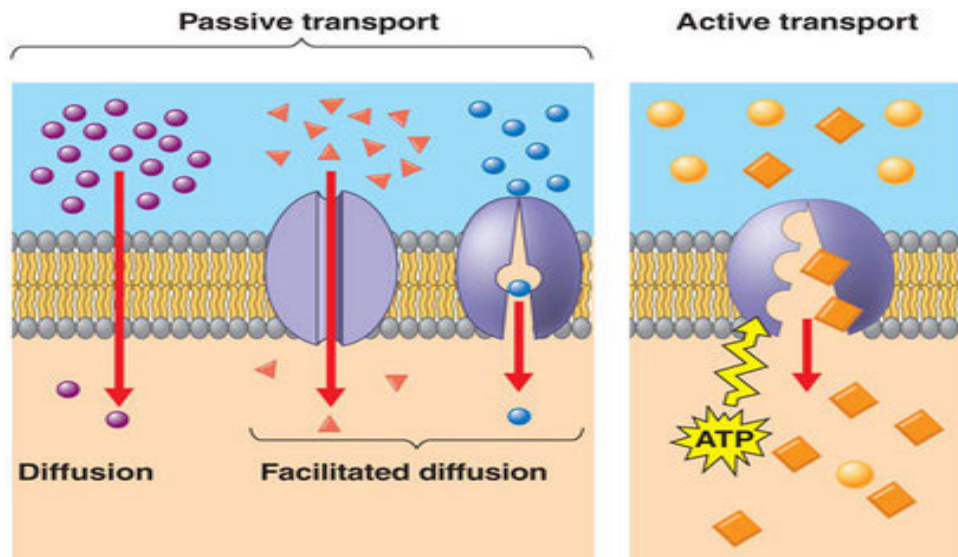
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*Transport function* is one of the most important functions of biological membranes. The most vital activity processes, such as absorption, excretion, propagation of nerve impulse, muscle contraction, ATP synthesis, etc. are bound up with movement of substances through membranes. Schematically, types of membrane transport can be represented as follows:



**Fig. 5.1b.** Types of membrane transport

**Exercise 5.2b.** Define the following terms and write down the answers.

*Passive transport* is \_\_\_\_\_  
 \_\_\_\_\_

*Active transport* is \_\_\_\_\_  
 \_\_\_\_\_

There are following types of passive transport: \_\_\_\_\_  
 \_\_\_\_\_

*Simple diffusion* is \_\_\_\_\_

*Osmosis* is \_\_\_\_\_

*Transport of substances through the membrane channels* is \_\_\_\_\_  
 \_\_\_\_\_

*Facilitated diffusion* is \_\_\_\_\_  
 \_\_\_\_\_

There are following types of active transport: \_\_\_\_\_  
 \_\_\_\_\_

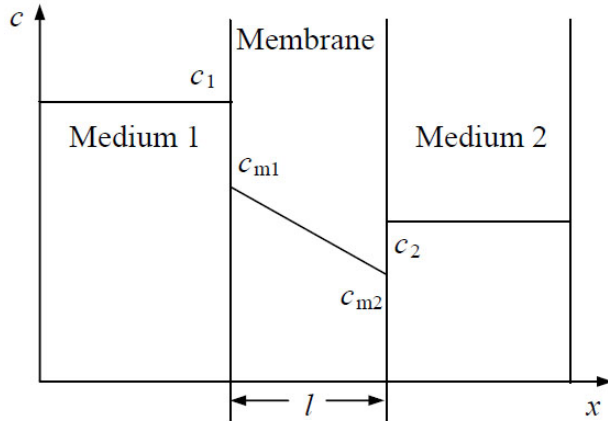
*Ion channels* are \_\_\_\_\_  
 \_\_\_\_\_

### **Passive Transport Across the Membrane**

They say that there is a *concentration gradient* if in two different regions of space the concentrations of the same substance are not equal. In this case, we will observe the phenomenon of diffusion or *passive transport* of the substance from an area of higher concentration to an area with lower concentration. This phenomenon will occur spontaneously (without energy consumption) until concentration of the substance in the two regions become the same and the total flow of the substance is zero. In a case of living cells such equilibrium may not be achieved if substances are continuously

synthesized or are spent in a course of chemical reactions in some part of space Passive transport of a substance along an axis is described by the *Fick's Law*:

$$J = -D \frac{dc}{dx}, \quad (5.1b)$$



**Fig. 5.2b.** Schematic representation of diffusion of substances between the cell and the external environment

where  $J$  is the *substance flux*. The flux is numerically equal to the quantity  $M$  of a substance transported per unit time  $t$  through a unit surface  $A$  perpendicular to the transport direction. Its units of measurement are  $[J] = \text{mol} / (\text{m}^2 \cdot \text{s})$ ;

$D$  is the *diffusion coefficient*. Its units of measurement are  $[D] = \text{m}^2/\text{s}$ ;

$dc/dx$  is the gradient of concentration in the  $x$  direction. The sign “-“ means that the flux is directed towards a low concentration area, i.e. results in decrease in the concentration gradient.

When considering the passive transport of uncharged particles through the membrane we can take gradient of concentrations as constant  $\frac{dc}{dx} = \frac{c_o^m - c_i^m}{l}$ , where  $l$  is thickness of the membrane.

Substituting this expression into the Fick's Law (5.1b) we obtain:

$$J = -D \frac{c_o^m - c_i^m}{l} = -DK \frac{c_o - c_i}{l}. \quad (5.2b)$$

Let's introduce a concept of a coefficient of distribution of the substance between the membrane and the ambient medium:

$$K = \frac{c_o^m}{c_o} = \frac{c_i^m}{c_i}. \quad (5.3b)$$

where  $c_o$  and  $c_i$  are concentrations of substance inside and outside the cell, respectively. The distribution coefficient  $K$  has no units of measurement. *Permeability coefficient* is defined as:

$$P = \frac{DK}{l}. \quad (5.4b)$$

**Problem 5.3b.** A flat bilayer membrane of thickness  $l = 10 \text{ nm}$  divides a chamber into two parts in which there is a substance of concentrations  $c_1 = 2 \text{ mmol L}^{-1}$  and  $c_2 = 30 \text{ mmol L}^{-1}$ . The substance flux through the membrane is  $J = 0.8 \times 10^{-3} \text{ mmol m}^{-2} \text{ s}^{-1}$ . Calculate the diffusion coefficient  $D$  of this substance if it is known that its distribution coefficient is  $K = 0.05$ .

**Data:**

**Solution:**

**Answer:** \_\_\_\_\_

**Problem 5.4b.** Calculate the coefficient of permeability for the substance which flow through the membrane is  $J = 5 \times 10^{-5} \text{ mol}/(\text{m}^2 \cdot \text{s})$ . The concentration of the substance inside the cell is  $1.8 \times 10^{-4} \text{ mmol} \cdot \text{L}^{-1}$  and outside the cell is  $3 \times 10^{-5} \text{ mmol} \cdot \text{L}^{-1}$ .

**Data:**

**Solution:**

**Answer:** \_\_\_\_\_

### Nernst Potential

In a living organism, there are ion solutions on both sides of a membrane. The concentrations of the same ion on both sides of the membrane often differ from each other. One of reasons of existence and maintenance of these concentration gradients is different permeability of the membrane to different ions. In this case the concentration gradient of a substance can be maintained by potential difference across the membrane (the so-called *transmembrane potential difference*). The value of *transmembrane potential difference* can be obtained by *Nernst equation*:

$$\Delta\varphi = \varphi_i - \varphi_o = \frac{RT}{zF} \ln \frac{c_o}{c_i}, \quad (5.5b)$$

where  $\varphi$  is the electrical potential measured in Volts;

$T$  is thermodynamic temperature measured in Kelvin degrees;

$z$  is the integer charge of the ion:  $\text{K}^+$ ;  $\text{Na}^+$ ;  $\text{Cl}^-$ ;  $\text{Ca}^{2+}$ ;

$c_o$  and  $c_i$  are concentrations of substance inside and outside the cell, respectively;

$F = 9.65 \times 10^4 \text{ C/mol}$  is Faraday's constant;  $R = 8.314 \text{ J}/(\text{K} \cdot \text{mol})$  is the molar gas constant.

**Problem 5.5b.** A semipermeable membrane divides two solutions of positive univalent ions of concentrations  $c_1 = 50 \text{ mmol}\cdot\text{L}^{-1}$  and  $c_2 = 500 \text{ mmol}\cdot\text{L}^{-1}$ . Find a potential difference across the membrane at a temperature of  $t = 37 \text{ }^\circ\text{C}$ .

Data:

Solution:

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*Answer:* \_\_\_\_\_

**Problem 5.6b.** Two solutions of lithium ions are divided by a flat bilayer membrane. Calculate ratio of ion concentrations in the solutions at which the equilibrium potential difference is  $\Delta\phi = 116 \text{ mV}$ . The medium temperature is  $t = 20 \text{ }^\circ\text{C}$ .

Data:

Solution:

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*Answer:* \_\_\_\_\_

**Problem 5.7b.** How many times an extracellular concentration  $c_o$  of sodium ions must exceed their intracellular concentration  $c_i$  in order to set Nernst potential to  $\Delta\phi = +50 \text{ mV}$  at a temperature of  $t = 27 \text{ }^\circ\text{C}$ .

Data:

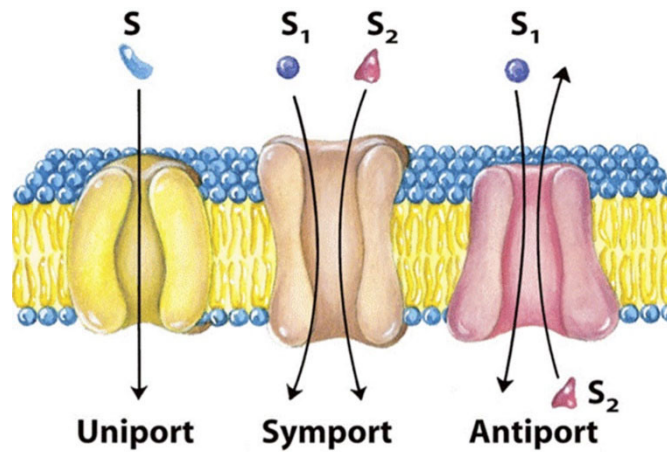
Solution:

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*Answer:* \_\_\_\_\_



## Active Transport Through the Membrane



**Fig. 5.3b.** Secondary active transport.

**Exercise 5.8b.** Complete definitions. Help yourself with picture.

*Secondary active transport* is \_\_\_\_\_

*Uniport* transports \_\_\_\_\_

*Cotransport* is \_\_\_\_\_

*Symport* transports \_\_\_\_\_

*Antiport* transports \_\_\_\_\_

**Exercise 5.9b.** Match the type of transport and the way in which it occurs:

Primary active transport

using any ion electrochemical potential gradient energy

Secondary active transport

performed using the energy of ATP

**Control questions**

1. Liposomes and their prospects for use in pharmacy.
2. How do you understand the concept of a concentration gradient? Describe in your own words.
3. What process is described by the Nernst equation?
4. What is the difference between the active transport of substances from passive?
5. What states are called the liquid-crystal and gel states of the membrane?

### ***Individual assignments***

1. A biomembrane of thickness 8 nm divides a compartment into two parts in which there is a substance of concentrations  $c_1 = 4 \text{ mmol L}^{-1}$  and  $c_2 = 35 \text{ mmol L}^{-1}$ . The substance flux through the membrane equals to  $1.2 \times 10^{-3} \text{ mmol m}^{-2} \text{ s}^{-1}$ . Calculate the diffusion coefficient of this substance if  $K = 0.16$ .
2. The concentration of the substance inside the cell is  $26 \text{ mmol L}^{-1}$ ; its concentration outside the cells is  $3.2 \text{ mmol L}^{-1}$ . The substance flux through a membrane which thickness is 9.6 nm, is  $J = 1.48 \times 10^{-3} \text{ mmol m}^{-2} \text{ s}^{-1}$ . Calculate the diffusion coefficient of this substance if it is known that its distribution coefficient is 0.33.
3. Calculate the distribution coefficient  $K$  of the substance through a membrane if the membrane thickness is  $8 \mu\text{m}$ , the diffusion coefficient of a substance is  $D = 65 \times 10^{-9} \text{ m}^2/\text{s}$ , the permeability coefficient of the substance is  $P = 1.17 \text{ m/s}$ .
4. The membrane thickness is 8.75 nm, the diffusion coefficient of a substance is  $90.12 \times 10^{-9} \text{ m}^2/\text{s}$ , the permeability coefficient of the substance is  $P = 10 \text{ m/s}$ . Calculate the distribution  $K$  of the substance.
5. A semipermeable membrane divides two solutions of univalent ions of concentrations  $c_1 = 30 \text{ mmol L}^{-1}$  and  $c_2 = 340 \text{ mmol L}^{-1}$ . Find a potential difference across the membrane at a temperature of  $t = 32 \text{ }^\circ\text{C}$ .
6. Two solutions of lithium ions are divided by a flat bilayer membrane. Calculate ratio of ion concentrations in the solutions at which the equilibrium potential difference is  $\Delta\phi = 96 \text{ mV}$ . The medium temperature is  $t = 26 \text{ }^\circ\text{C}$ .
7. How many times an extracellular concentration  $c_o$  of sodium ions must exceed their intracellular concentration  $c_i$  in order to set Nernst potential to  $\Delta\phi = +62 \text{ mV}$  at a temperature of  $t = 30 \text{ }^\circ\text{C}$ .