

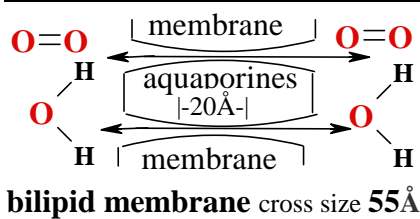
The **Water channels**, allow the passive but selective movement of **Water** and **O₂,NO,CO** across **cell wall** and **subcellular membranes** like as mitochondria, endoplasmic reticulum, peroxisomes, Golgi, lysosomes.... . **Aquaporins** have been classified into **two** sub-families:

I) strict **Aquaporins** that only allow the passage of **Water, O₂, NO, CO** and II) the less selective **aquaglyceroporins** that **transport Water** and other neutral **solutes**, such as **Glycerol, CO₂** or **urea**.

Recently, the identification and characterization of a number of archaeal and bacterial **Aquaporins** suggested the existence of a **third** sub-family; one that is neither a strict **Aquaporin** nor an **aquaglyceroporin**. The function and phylogeny of this **third** family is still a matter of debate.

Water channels H₂O common O₂, NO, CO: an overview

AQP0	+ Cl⁻, NO₃⁻ eye-lens cells; thin junctions between fiber cells AQP0 with a measured Water permeability <u>15-fold lower</u> than that of AQP1 at pH 6.5 ; AQP0 is <u>reduced</u> a further <u>three fold</u> at pH 7.5 AQP0 induce a gating effect <u>close</u> conformations of <u>extracellular loop A Met176.His40</u> AQP0 becomes more constrained near the conserved Ar/R constriction site
AQP1-	+ Cl⁻, NO₃⁻ , Aquaglyceroporins: red blood cell (RBC) , <u>apical & basolateral membranes</u> of epithelial brain cell, rodent brain cell AQP1-null humans kidney proximal-tubule water reabsorption gastrointestinal tract Water absorption in the teleost intestine the ovary and in the oocyte ; salivary gland ;
AQP2	urinary bladder granular kidney cells & subcellular vasopressin regulated urine concentration (25% of the blood filtrate) trans located from the <u>cytoplasmic pool</u> to the <u>apical plasma membrane</u> of the granular <u>cells</u> of the pelvic patch and urinary bladder
AQP3	+ <u>Aqua glyceroporins</u> , urea: gastrointestinal tract Water absorption; rodent brain cell astrocyte end-feet Water enters in the principal <u>cell</u> through AQP2 and exits through located in the basolateral membranes trachea basal AQP3 & ciliated columnar AQP4 <u>cells</u>
AQP4	Rodent-brain ;basolateral membrane of ciliated columnar <u>cells</u> alveolar epithelium;salivary gland
AQP5	stomach, duodenum, pancreas, airways, lungs, salivary gland, sweat glands, eyes, lacrimal glands, and the inner ear tears & pulmonary sub mucosal glands secretions <u>apical membrane & rodent brain cells</u>
AQP6	+ Cl⁻, NO₃⁻ multi permeable channel ;lens cells; may <u>play a role</u> in the body <u>acid-base homeostasis</u> in the intracellular vesicles of <u>acid-secreting intercalated cells</u> of the RCD colocalized with the H⁺-ATPase be Hg²⁺-inhibit able Water channel function is activated by Hg²⁺ and low pH
AQP7	+ <u>Aquaglyceroporins</u> , urea; kidney proximal tubule epithelium cell glycerol reabsorption ; together with AQP1 <u>in the brush border</u> in the concentration of urine taking place in the proximal nephron 75% of the blood filtrate which is 150–180 L per day
AQP8	NH₄⁺ ;lens & kidney intracellular proximal tubule & small intestine absorptive:epithelium cell in the concentration of urine taking place in the proximal nephron also in mitochondria 75% of the blood filtrate which is 150–180 L per day & rodent brain cell
AQP9	+ <u>Aquaglyceroporins</u> , urea purines, pyrimidines & monocarboxylates, arsenite ; <u>apical membrane</u> of brain & small intestine absorptive epithelial & rodent brain & glial cells
AQP10	+ <u>Aquaglyceroporins</u> , urea ; small intestine absorptive epithelial <u>cells</u>
AQP11	“super aquaporins ” or sub <u>cellular</u> ; kidney cytoplasm of the proximal tubule & rodent brain cells
AQP12	“super aquaporins ” or sub <u>cellular</u> H₂O Channel is roughly 20-Å long and has a diameter 1.1 Å. Water



channel proteins (WCPSs) are **trans membrane proteins** that have a specific three-dimensional structure with a **pore** the **SF radius 1.1 Å** is close average to **radius of water H-O-H** longitudinal **1.4 Å** and **0.55 Å** bent size of dipole.

It can be permeated by **Water & O₂, NO, CO** molecules as solutes.

Aquaporins are large families (over **450 members**) that are present in all kingdoms of life. **Water** permeability, allowing permeation of **3 × 10⁹ water** molecules per **monomer** per second AQP1 and

other, which strictly prevents the conduction of protons **H⁺**.

Phosphorylation to trigger the **membrane trafficking** of AQP1, AQP2, AQP5, and AQP8, and the gating of AQP4.

Cation conductance has been induced in AQP1 by activation of cyclic GMP-dependent pathways and

<http://aris.gusc.lv/ChemFiles/Aquaporins/WCPsAQPsIUBMBLife09/AQP0-11.doc> was blocked by **Hg²⁺**

Red blood cells against colligative osmomolar concentration in water solutions

Water and oxygen osmosis against osmo molar concentration gradient crosses cell membranes

Osmosis is organised for **H₂O** and **O₂** movement against concentration gradients-difference of colligative properties $\Delta C_{osm} = i\Delta C_M$ through an **Aquaporins** across cell **membranes** to form the osmotic pressure:

$$\pi = i\Delta C_M RT \text{ (kPa) ,}$$

where $R=8,3144 \text{ J/(mol}\cdot\text{K)}$ universal gas constant,

T temperature in Kelvin's degree (K) $T=t+273.15$ (if $t=37^\circ$ than $T=37+273.15=310.15 \text{ K}$).

Note: Transfer **water** and oxygen molecules through membrane aquaporin tunnel in erythrocytes with rate $3\cdot 10^9 \text{ sec}^{-1}$ in both directions transfer 3000 oxygen molecules in one second.

Mechanism of osmosis through membrane aquaporins drive colligative concentration gradient

Na⁺Cl⁻ => **Na⁺+Cl⁻** $m=2$ electrolyte dissociation $\alpha=1$ the concentration gradient doubled as i is 2

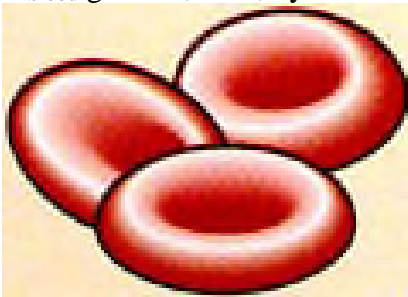
$i=1+\alpha(m-1)=1+1(2-1)=2$; $i\Delta C_M = 2\Delta C_M = \Delta C_{osm}$ and pressure on membrane is $\pi = 2\Delta C_M RT = \Delta C_{osm} RT$.

Water H₂O, O₂ oxygen flow left side against the concentration gradient from 0 to $C_{osm}=0.305 \text{ M}$ because **Na⁺Cl⁻** ions make osmo molar concentration left side $C_{left}-C_{right}=C_{osm}-0=C_{osm}=iC_M$ and **close H₂O, O₂** flow to right side.

$$C_{blood} = C_{osm} = i_1C_1 + i_2C_2 + i_3C_3 + \dots = \sum i_k C_k = 0,305 \text{ M}$$

Human erythrocytes red blood cells with osmo molar concentration 0.305 M of all solutes sum $\sum i_k C_k$:

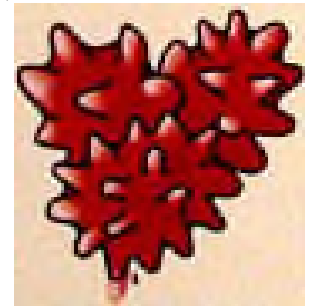
C_{blood} glucose, salts, hydroxonium **H₃O⁺**, hydroxyl **OH⁻** ions, amino acids, proteins, bicarbonate etc.



Isotonic medium
 $C_{blood} = 0.305 \text{ M}$

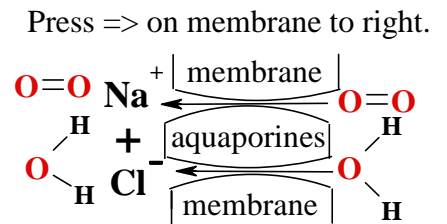


Hypotonic medium
distilled water 0 M
or at least



Hypertonic solution
 $C_{Hyperton} \geq 0,4 \text{ M}$.
Hypertonic salt solutions to apply for purulent wounds, because pumps **water** toxic compounds out and stimulates **blood** circulation.

osmo molar concentration $C_{Hypoton} \leq 0,2 \text{ M}$.
Hypotonic **water** medium the flow is greater towards the cell against the concentration gradient-difference $0.305 - 0,2 = 0,105 \text{ M}$ and the cell puffs up until its membrane is broken but content leak in plasma.



Osmosis **H₂O** and **O₂** against concentration gradient through alveolar epithelial membrane

A) Oxygens O₂ from **AIR** 20.95% **O₂** ↑ gas assimilation reaction dissolution in water to form **O₂aq** exothermic $\Delta H_r = -55,7 \text{ kJ/mol}$ and exoergic $\Delta G_r = -27,7 \text{ kJ/mol}$ as water soluble oxygen :

1) $O_{2AIR} + H_2O \rightleftharpoons H_2O + O_{2aq} + Q + \Delta G$. Penetrate in Human body through aquaporins by concentration gradient from $[O_2] = 9,768 \cdot 10^{-5} \text{ M}$ to **venous** blood $[O_{2aq}] = 1,85 \cdot 10^{-5} \text{ M}$.

2) $\Delta G_{O_2} = RT \ln([O_{2Blood}]/[O_{2aq}]) = -4,29 \text{ kJ/mol}$ exoergic entrance human organism;

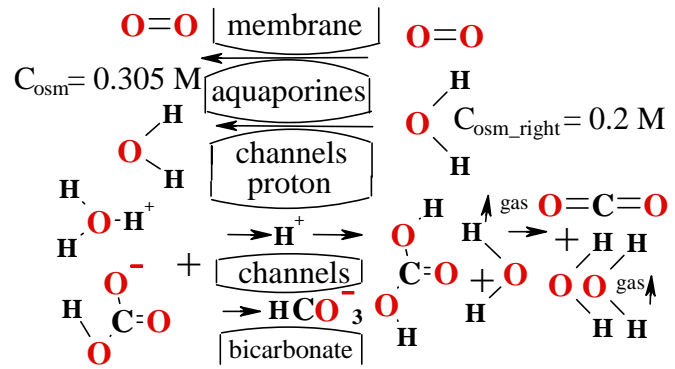
3) $O_{2aq} + H_2O \xrightarrow{\text{Aquaporins}} H_2O + O_{2aq} + \Delta G$ against concentration gradient 0,305 M / 0,2 M:

$$\Delta G_{H_2O} = RT \ln([H_2O]_{right}/[H_2O]_{left}) = -8,3144 \cdot 310,15 \cdot \ln(0,305/0,2) = -1.088 \text{ kJ/mol}$$

exoergic $\Delta G_{O_2} = -5,379 \text{ kJ/mol}$. **Deoxy** hemoglobin **Hb_T** adsorbs 4 **O₂aq** from blood plasma of inspired fresh **AIR** releases four protons **4H⁺** and 4 **HCO₃⁻** stabilizing arterial $[O_2] = 6 \cdot 10^{-5} \text{ M}$ concentration **4O₂aq** + $(H^+His63,58)_4 Hb_T \rightleftharpoons Hb_R(O_2)_4 + 4H^+$.

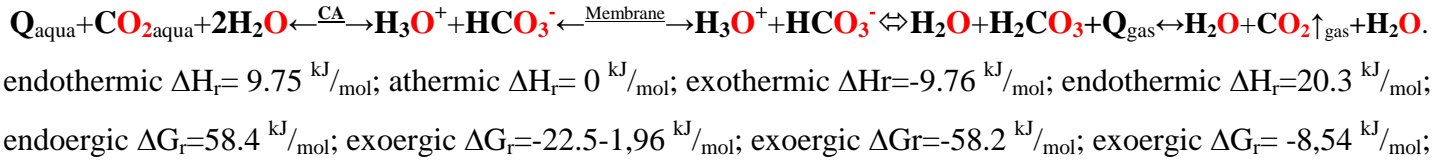
$$\text{Total exothermic } \Delta H_r = -55,7 \text{ kJ/mol and exoergic } \Delta G_{O_2} = -27,7 + -4,29 + -1.088 = -33.078 \text{ kJ/mol}$$

Osmosis is **water** and oxygen flow left side against gradient of concentration 0.2 M to $C_{osm}=0.305$ M because water and oxygen flow to right side closed by made left side osmo molar $C_{left}-C_{right}=C_{osm}-C_{osm_right}=\Delta C_{osm}$ concentration as difference $\Delta C_{osm}=0.105$ M. $C_{osm}=i_1C_1 + i_2C_2 + i_3C_3+\dots=\Sigma i_kC_k=0,305$ M; $C_{osm_right}=0,2$ M; $\Delta C_{osm}=0.305-0.2=0.105$ M



Breath out **H₂O** , **CO₂** in endothermic but exoergic reactions on alveolar epithelial surface

B)



endothermic $\Delta H_r = 9.75$ kJ/mol; endothermic $\Delta H_r = 54,5$ kJ/mol; summary endothermic $\Delta H_r = 64,25$ kJ/mol; endoergic $\Delta G_r = 58.4$ kJ/mol; exoergic $\Delta G_r = -82,1$ kJ/mol; summary exoergic $\Delta G_r = -23,7$ kJ/mol;

Venous **deoxy Hb_T shuttle** adsorbs four **oxygen** 4O₂ molecules, acidify water medium with 4H⁺, promoting **CO₂** breathe out as increase production of H⁺ , **HCO₃⁻** $459 \cdot 6 \cdot 10^{-5} M = 0,0275 M = [HCO_3^-]$ amounts shifts equilibrium to right $H^+ + HCO_3^- + Q \leftrightarrow H_2O + CO_2 \uparrow_{gas}$ via membrane channels. So pH=7,36 remains constant, as bicarbonate ion and hydrogen ion produce **CO₂** right side.

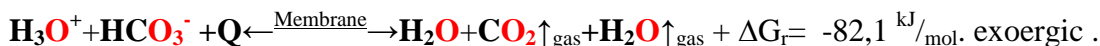
The epithelial cell surface of **lungs** has the specific building. $S = 950 \text{ nm} \times 950 \text{ nm} = 0.9 \mu m^2$ is surface area with super thin 0.6 nm **water** layer volume: $0.5415 \cdot 10^{-3} \mu m^3 = 0.5415 \cdot 10^{-18} L$. Created acidity in thin **water** layer volume increases up to pH=5.5 if one proton H⁺ crosses the membrane channels reaching the surface. Hydrogen ion concentration is: $[H_3O^+] = 10^{-pH} = 10^{-5.5} M$. Respiration in **lungs** Hemoglobin released protons H⁺ during oxygen adsorbtion for total amount concentration:

$[O_{2,adsorbed}] = [H_3O^+] = 459 \cdot 6 \cdot 10^{-5} M = 0,02754 M$ forms hydrogen ion concentration gradient:

$[H_3O^+]_{right} / [H_3O^+]_{left} = 10^{-5.5} / 0,0275$, which drives exoergic $\Delta G = -22,5$ kJ/mol proton movement through epithelial cell membrane proton channels: $H_3O^+_{left} \xleftarrow{\text{proton channel}} H_3O^+_{right} + \Delta G$. General process

$H_2O + CO_2 \uparrow_{gas} + H_2O \uparrow_{gas}$ require heat supply endothermic $\Delta H = 54,5$ kJ/mol to drive spontaneous

$\Delta G = -82,0679$ kJ/mol products evaporation **CO₂** \uparrow_{gas} and **H₂O** \uparrow_{gas} keeping moisture **H₂O** on surface of membrane. Hydrogen ions water acidity shift endothermic $\Delta H_r = +54,5$ kJ/mol and exoergic $\Delta G_r = -82,1$ kJ/mol decomposition **H₃O⁺ + HCO₃⁻** breath out to AIR **CO₂** \uparrow_{gas} with **H₂O** \uparrow_{gas} :



Aquaporins are wide class of **membrane crossing channel** proteins, **which are integrated** in all living organisms: **animals, plants, bacteria**. On Cell membranes effecting Physiology, Biochemistry and Health. **Aquaporins** are large families (over 450 members) that are present in all kingdoms of life.