

# Resting membrane potential

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## Definition:

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it is the potential difference between **inside** and **outside** the cell membrane with inside **relatively negative** to outside.

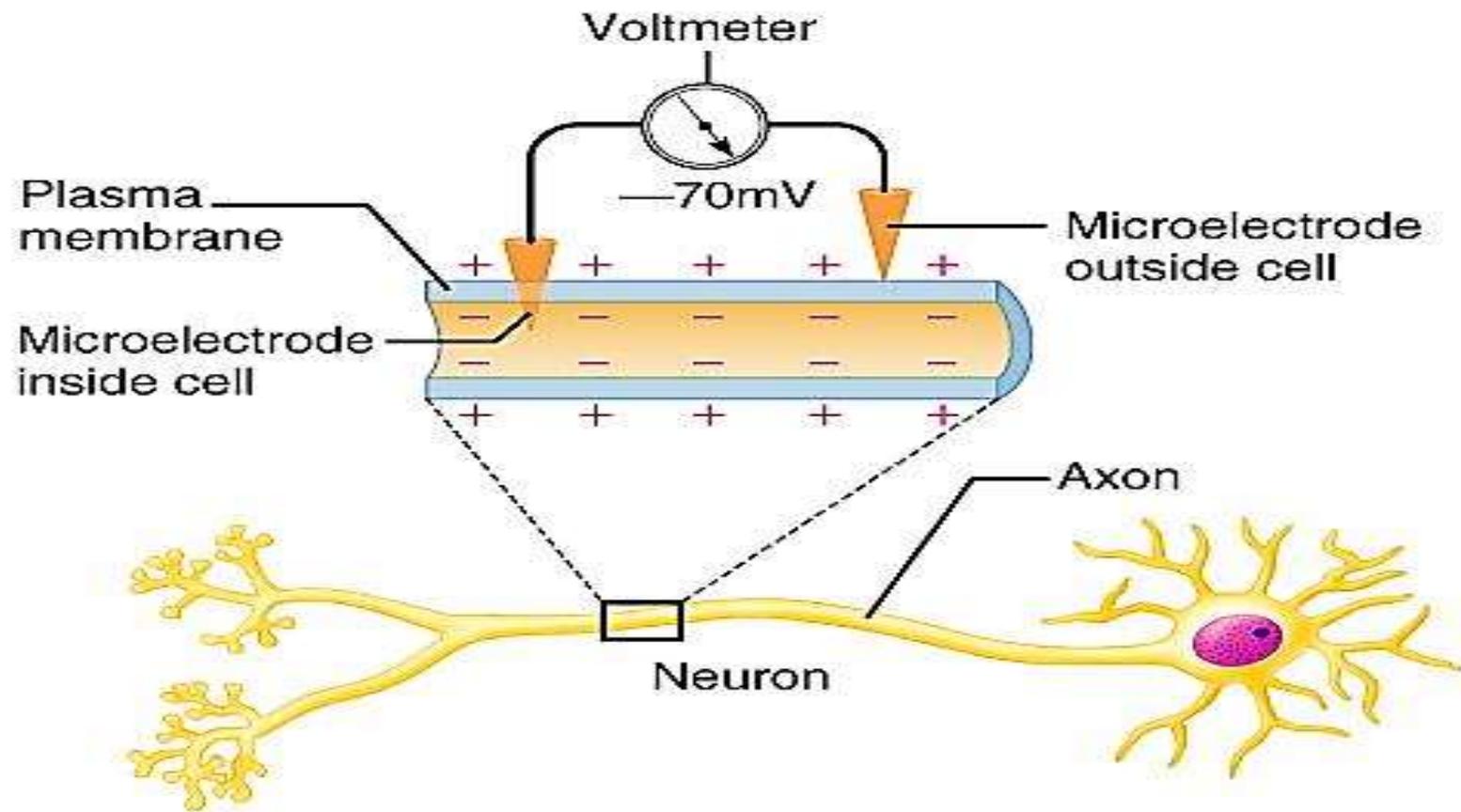
It ranges from **-9** mvolt to **-100** mvolt .

in RBCs it equals **-30** mvolt.

in medium sized nerve it equals **-70** mvolt

in large sized nerve it equals **-90** mvolt

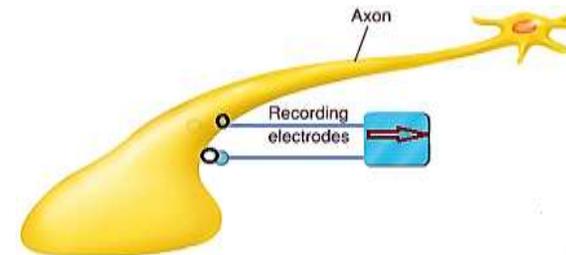
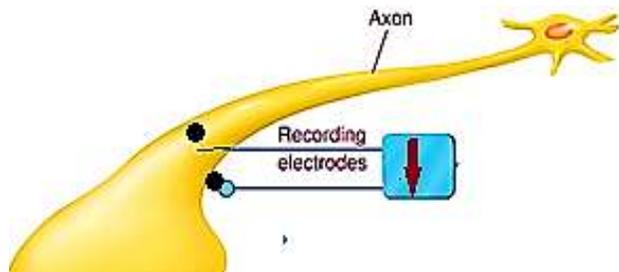
in the muscle **-90** mvolt.



## Evidence:

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- ❖ If **2 electrodes** are put on the outer surface of the cell membrane or on the inner surface of cell membrane, no deflection on the galvanometer **(there is no potential difference)**.
- ❖ If one electrode is put **on outside** and the other **inside** the cell membrane there is deflection in the galvanometer.



## Causes of RMP

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It is due to **unequal distribution** of ions across the cell membrane

**outside:**  $\text{Na}^+$ (142mEq./L),  $\text{Cl}^-$ (103mEq./L),  $\text{HCO}_3^-$ (28mEq./L)

**inside:**  $\text{K}^+$ (140mEq./L) & protein<sup>-</sup> (40mEq./L))

more **cations outside**

and more **anions inside.**

This **unequal distribution** is caused by three factors:

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1- Selective permeability of the cell membrane

2- Na<sup>+</sup> - K<sup>+</sup> pump

3- The membrane is impermeable to the intracellular protein anions with large molecular weight (protein)

# [I] Selective permeability of the cell membrane:

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The cell membrane is **semipermeable** and has pores (channels)

**These channels are:**

- 1- leakage channels (passive)
- 2- gated channels (active)

# 1- leakage channels (passive)

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- Watery pathway through protein molecules.
- Open all time.
- Not gated
- Tube shaped, 70 (Angstrom) in diameter.
- Highly selective and this **selectivity depend on**
  - Size of channel.
  - Charges inside the channel.
  - Size of molecule.
  - Charges of molecules.
  - Shape of channel.
  - Concentration gradient of molecules.

**N.B:** *Na<sup>+</sup> channels have negative charge while K<sup>+</sup> channels have no charge*

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*To which ion the cell membrane is more permeable  
( $\text{Na}^+$  or  $\text{K}^+$ ) and why?*

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The membrane is more permeable to **K<sup>+</sup>** than to **Na<sup>+</sup>** ions because the hydrated **K<sup>+</sup>** is smaller in size than the hydrated **Na<sup>+</sup>**. So, **K<sup>+</sup>** permeability **50-100 times** greater than Na<sup>+</sup> permeability.

## 2-Gated channels (active)

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- have gates

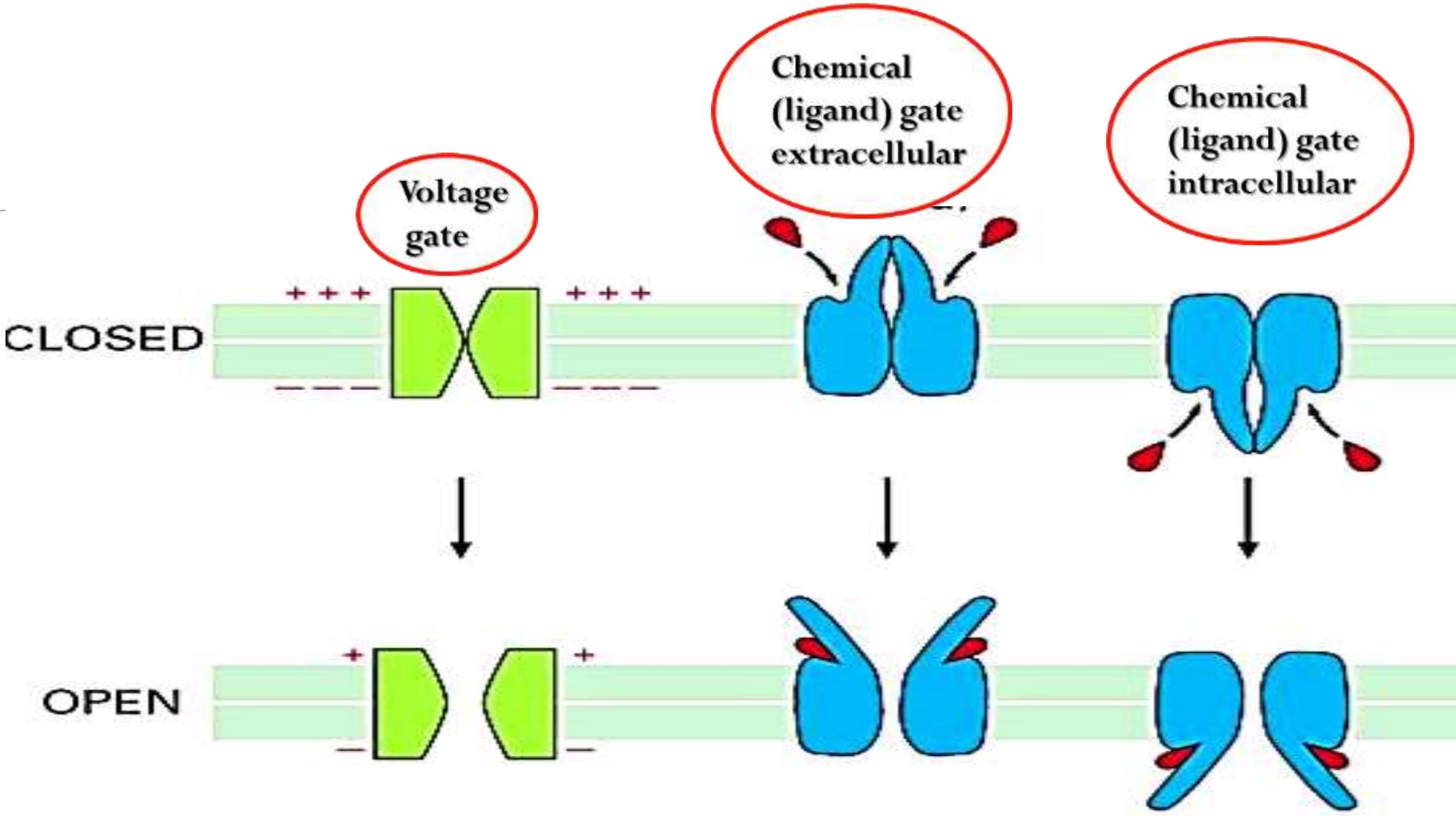
These gates may be opened or closed by changing the shape of protein molecules in the channel in response to various agents:

1- if change in **ionic composition** (Voltage gated channels)

2- if binding **with certain chemical substances** (ligand gated channels).

\*\*Na<sup>+</sup> channels have **outer** activation and **inner** inactivation gates .

\*\* K<sup>+</sup> channels have one inner gate.



## **N.B.**

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During rest the movement of ions is through the leakage channels  
During stimulation and action potential it occurs via the gated ones.

# Membrane permeability:

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**As regard to Na<sup>+</sup>:** Na<sup>+</sup> tend to diffuse **into the cell** according to its **concentration** and **electrical** gradient until it is balanced by Na<sup>+</sup> efflux according to electrical gradient, at equilibrium the membrane potential = **+61 mv.**

**As regard to K<sup>+</sup>:** K<sup>+</sup> diffuse **from inside to outside** according to its **concentration** gradient till +ve charge outside repel more K<sup>+</sup> diffusion (electrical gradient). at equilibrium the membrane potential = **-94 mv.**

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## Nernst equation

- used to determine the **equilibrium potential** of each ion.
- Electromotive force (EMF) =  $\pm 61 \log (\text{concentration inside}/\text{conc. outside the membrane})$ .  
**e.g.** for Na<sup>+</sup> EMF =  $-61 \log (15/150) = +61$  millivolt.  
for K<sup>+</sup> EMF =  $-61 \log (150/5) = -94$  millivolt.
- According to the degree of the permeability of the membrane to Na<sup>+</sup> and K<sup>+</sup> the potential will be determined.
- **Goldman equation** is used to determine the equilibrium potential of all ions, which is about **-86** millivolt (near to the equilibrium potential of K<sup>+</sup> indicating that K<sup>+</sup> permeability is the main force responsible for the resting membrane potential).

## [II] Na<sup>+</sup> - K<sup>+</sup> pump:

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### At rest:

some Na<sup>+</sup> can enter inside the nerve fibre.

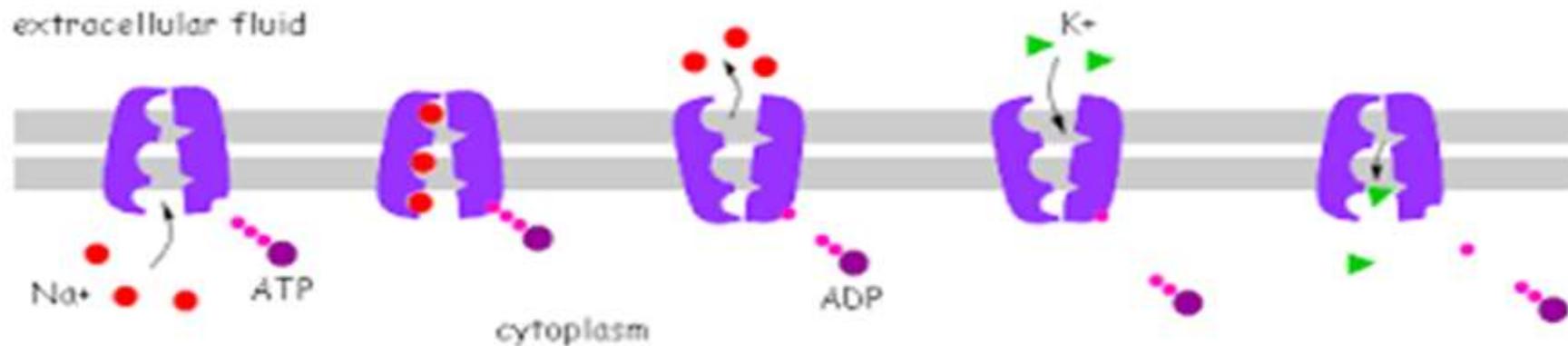
- **During action potential** large number of Na<sup>+</sup> ions enter the cell and K<sup>+</sup> efflux occur so,

the Na<sup>+</sup> pump is required **to return** Na<sup>+</sup> outside (against conc. and electrical gradient) and K<sup>+</sup> pump is required for return K<sup>+</sup> **inside** the cell against the concentration gradient only.

## Na<sup>+</sup>-K<sup>+</sup> pump needs:

- ✓ **energy source (ATP)**
- ✓ **ATPase enzyme** for liberation of energy.
- ✓ **large carrier protein** present in the cell wall.

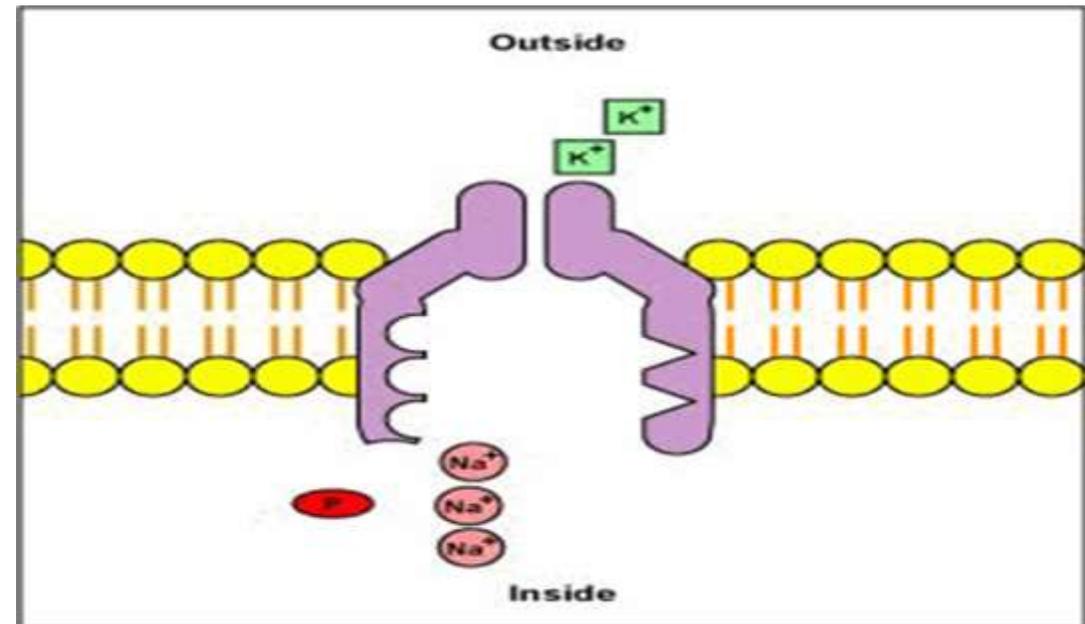
Its internal surface has **3 receptors** for Na<sup>+</sup> and ATPase,  
the external surface has **2 receptors** for K<sup>+</sup>.



## When activated :

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by energy from splitting of ATP by ATPase enzyme the pump rotates to push: 3  $\text{Na}^+$  to outside and 2  $\text{K}^+$  to inside the cell membrane.



## Importance of the Na<sup>+</sup>-K<sup>+</sup> pump

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- 1- Maintenance of Na<sup>+</sup> (extra cellular) and K<sup>+</sup> (intracellular).
- 2- It is an electrogenic pump as it causes **RMP** is more negative inside (- 4 mvolt), (2K<sup>+</sup> influx and 3 Na<sup>+</sup> outflux).
- 3- Control of cell volume as if Na<sup>+</sup> remains inside the cell, water enters by osmosis and the cell swells.

<b>Selective permeability</b>	<b>Na<sup>+</sup>-K<sup>+</sup> pump</b>
<p data-bbox="453 639 1098 793">-Initiation of the RMP(-86 mvolt).</p> <p data-bbox="453 886 1054 953">-<b>Passive</b> process.</p> <p data-bbox="453 1043 894 1196">-K<sup>+</sup> is mainly responsible</p>	<p data-bbox="1177 639 1862 793">-Maintenance of RMP by – 4 mvolt.</p> <p data-bbox="1177 903 1727 971">-<b>Active</b> process</p> <p data-bbox="1177 1061 1735 1213">-Na<sup>+</sup> and K<sup>+</sup> are responsible</p>

III] The membrane is impermeable to the intracellular protein  
**anions** with large molecular weight.

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So

**more negative charges inside the cell**

**(Donnan effect)**

This protein regulates diffusion of other anions and cations until  
reach the **Donnan's equilibrium**

$$\text{Cl}^- \times \text{K}^+ (\text{inside}) = \text{Cl}^- \times \text{K}^+ (\text{outside})$$

## **N.B.**

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- 1- the more the negativity of the resting membrane the more the excitability of the tissue
- 2- the most excitable tissue in the body are nerve and muscles.

Thank  
You

