

# BB 101

## MODULE: *PHYSICAL BIOLOGY*

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# Review of Lecture 2

- How surrounding medium affects functioning of biological systems?
- Critical Viscous Force and Reynolds number
- Life at low Reynolds number
- A low-Reynolds number microorganism can't swim by executing *geometrically reciprocal motion*
- Swimming of microorganism
- Swimming by reciprocal motion in non-Newtonian fluid

## Summary so far.....

We looked at the **forces** at molecular and cellular scales

Cellular world is predominately governed by **viscous forces**

As a consequence, **inertial forces** can be safely ignored in most cases

## What about energies?

# Thermal Energy and Thermal Forces

- Proteins and cells are subjected to thermal forces, arises due from **collision** of water and other molecules in their surrounding fluid
- These collision forces are called **thermal forces** because their magnitude is proportional to temperature of the fluid molecules.

# Thermal Motion and Thermal Energy

- The resulting movement of object is called **thermal motion**, and object is said to have **thermal energy**
- Since **thermal forces** are randomly directed, the resulting thermal motion is characterized by frequent changes in direction
- Diffusion of a free particle or object is called **Brownian motion**

# Brownian Motion



Figure Source: <http://www.nndb.com/people/050/000100747/>

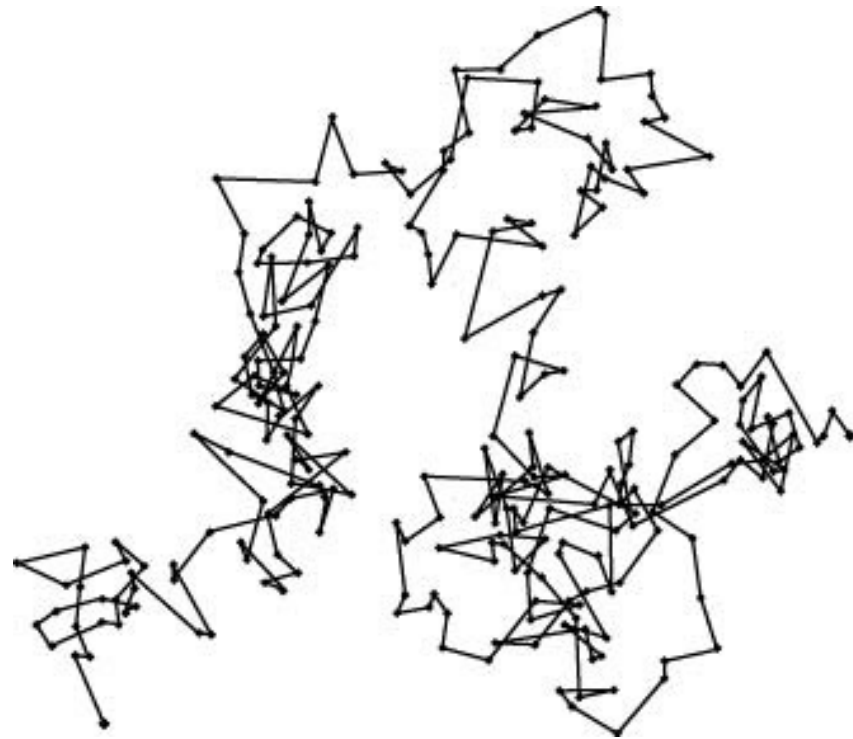
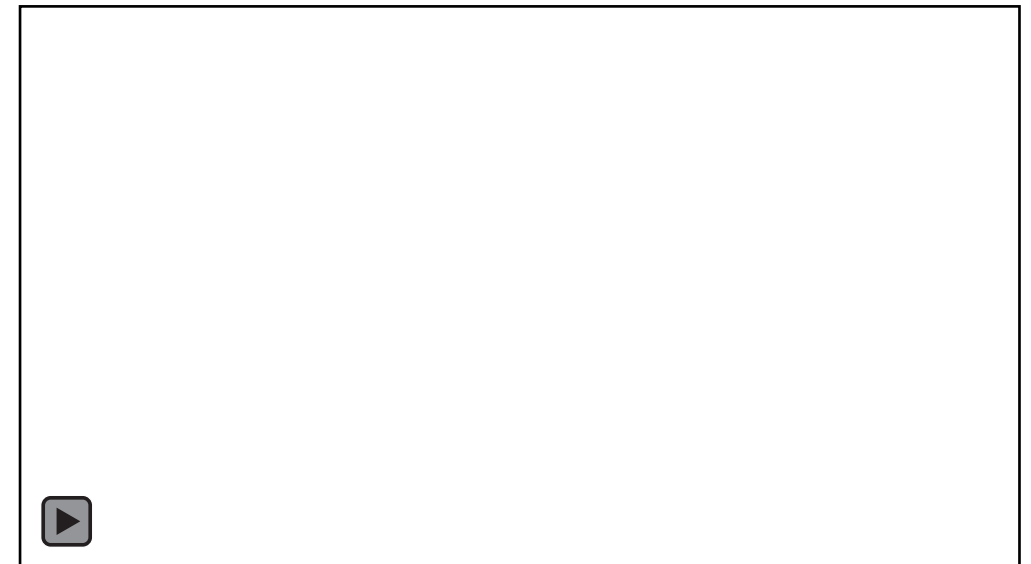


Figure Source: [http://www.doc.ic.ac.uk/~nd/surprise\\_95/journal/vol4/ykl/report.html](http://www.doc.ic.ac.uk/~nd/surprise_95/journal/vol4/ykl/report.html)

In 1828, botanist Robert Brown noticed that pollen grains suspended in water dance in zig-zag manner

- Initially thought that it was signature of life
- Careful observer and proceeded to check his assumption
- Repeated observations with many lifeless particles, and all of them showed the same (suspension of coal dust, pulverized stone etc.)



Watch Video of Brownian Motion of pollens in water  
<https://www.youtube.com/watch?v=R5t-oA796to>

# Thermal Energy

- We saw that objects suspended in fluid can gain thermal energy and this thermal energy can make them to dance
- The thermal energy at temperature  $T$  is given by  $k_B T$ , where  $k_B$  is Boltzmann constant
- Thermal energy at room temperature

$$k_B T \approx 4.14 \text{ pN nm } (T=300\text{K})$$

*Is this thermal energy important?*



# Relative Importance of Thermal Energy

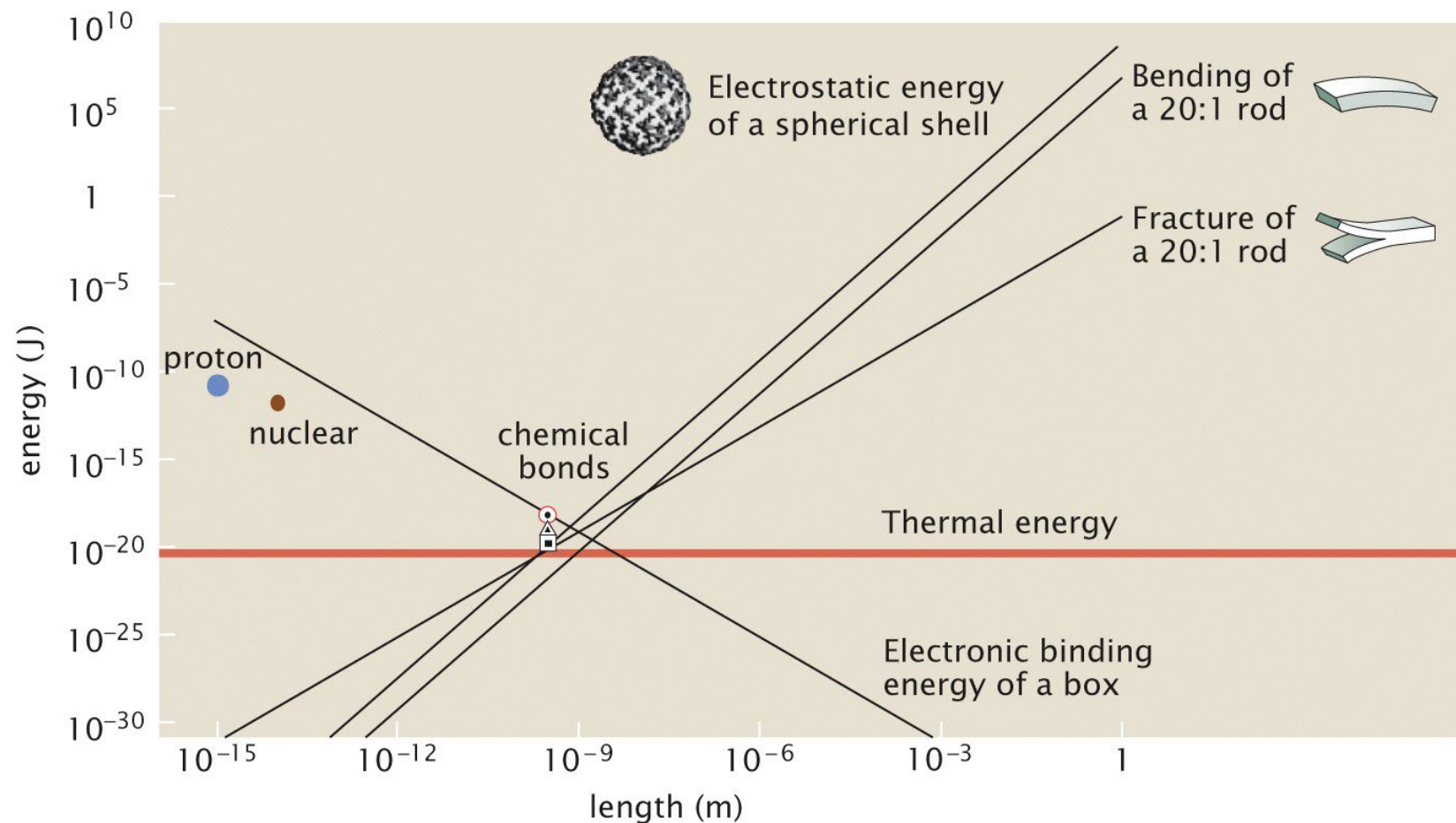


Figure 5.1 Physical Biology of the Cell, 2ed. (© Garland Science 2013)

- The electrostatic energy is estimated for a model spherical protein with polar residues on its surface and for which all of the polar residues are stripped of a single charge
- Binding energy is estimated by considering the effects of confining a free electron in a box of that length scale.

# Relative Importance of Thermal Energy

- At the scale of macromolecule of the cells (nm) deterministic energies of bonding, charge rearrangement and molecular rearrangement are comparable
- Because each of these forms of energy is of comparable scale and effectively interchangeable at the molecular level, a living organism that needs to generate motion, heat, electricity, and biomolecular synthesis is expert at energetic interconversions.

# Thermal Energy and Biological Systems

It becomes important to consider thermal energy at macromolecular scales

Thermal energy can be safely ignored at macroscopic level.

It turn out that state of a biological system at molecular scale is decided by the competition between deterministic energy and thermal energy

# Boltzmann's Law

Fundamental physical law that describes how probability of finding a molecule in a certain state depends on the energy of that state and surrounding temperature

A particle or molecule always tends to remain in its lowest energy state

At non-zero temperature, due to molecular collisions, they can spend their time in higher energy states

# Boltzmann's Law

Boltzmann's law says that if such a particle is in thermal equilibrium, then the probability  $p_i$  of finding the particle in state  $i$  that has energy  $U_i$  is given by

$$p_i = \frac{1}{Z} e^{-\frac{U_i}{k_B T}}$$

Where  $z = \sum_i e^{-\frac{U_i}{k_B T}}$  is called **partition function**

The exponential term is called **Boltzmann factor**

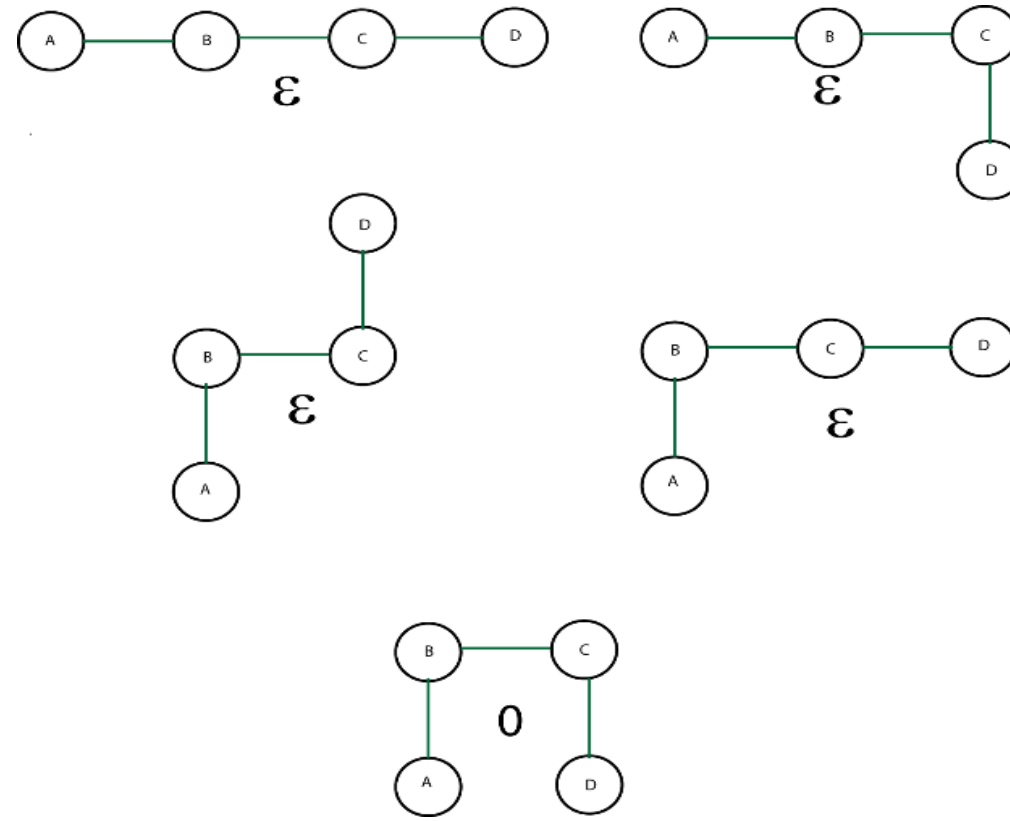
**Partition function tells you how the system will be partitioned or divided into different possible states**

# Partition Function

Consider a protein of four distinguishable amino acids with energy  $\epsilon$  kept in water. This protein switches between open conformation(s) and closed conformation(s) in a plane due to change of bond angles by  $90^\circ$  due to thermal fluctuations. Assume that energy of this system is not changed due to change of bond angles. However, if the distance two un-bonded amino acids due to change of bond angles become equal to bond length, then energy of the protein reduces by  $\epsilon$ . Find out the partition function ( $Z$ ) for this protein in the limit  $T \rightarrow 0$  and  $T \rightarrow \infty$ ?



# Partition Function



# Boltzmann's Law: Some comments

- Boltzmann's law is very general.
- The energy could correspond to particle's potential energy (gravitational, elastic or electrical), its kinetic energy or the energy associated with its phase, or electronic or chemical state
- If there are just two states with energies  $U_1$  and  $U_2$  (and energy difference  $\Delta U = U_2 - U_1$ ) then

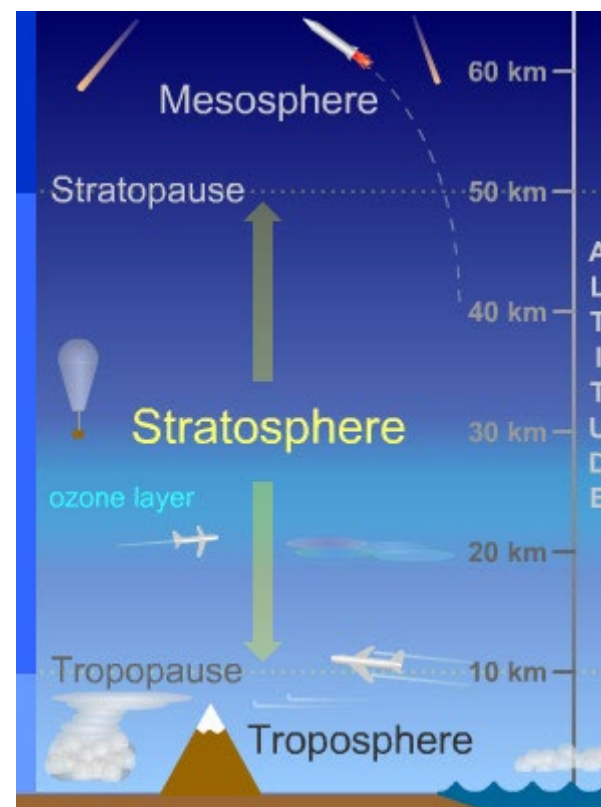
$$\frac{p_2}{p_1} = e^{-\frac{\Delta U}{k_B T}}$$



# Applications of Boltzmann's Law

## *Height of Atmosphere*

At highest point of atmosphere, the probability of finding oxygen molecule become  $1/e$  time that of ground level.



Height  $\sim 7.8$  km

# Applications of Boltzmann's Law

## *Nernst Equation*

Most spectacular application of biological electricity by cells is the action potential in the nerve cells

This is used to rapidly propagate information from the nerve cell body to the tip of axon

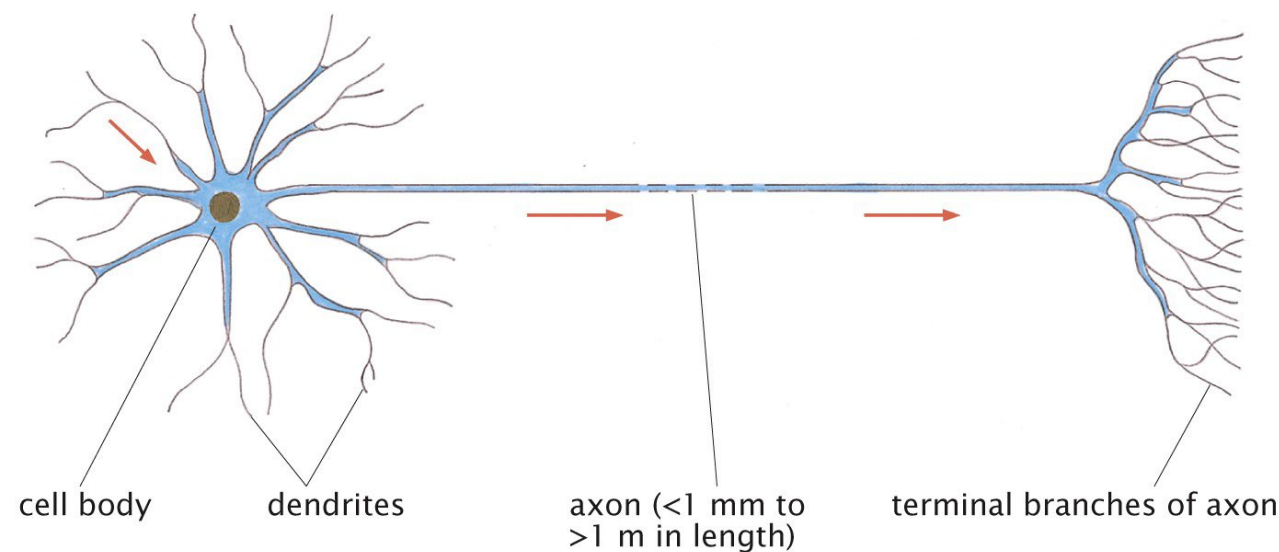


Figure 17.1 Physical Biology of the Cell, 2ed. (© Garland Science 2013)

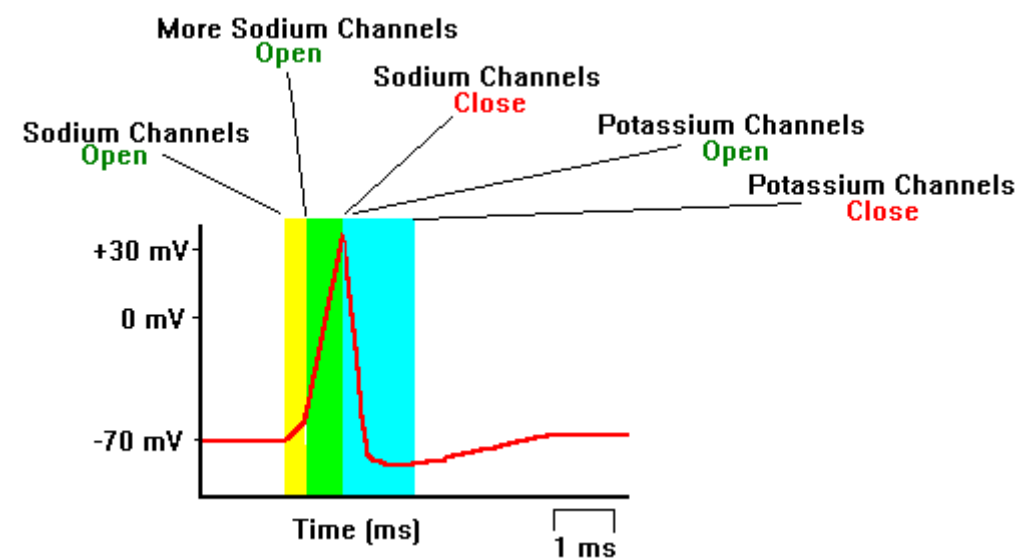
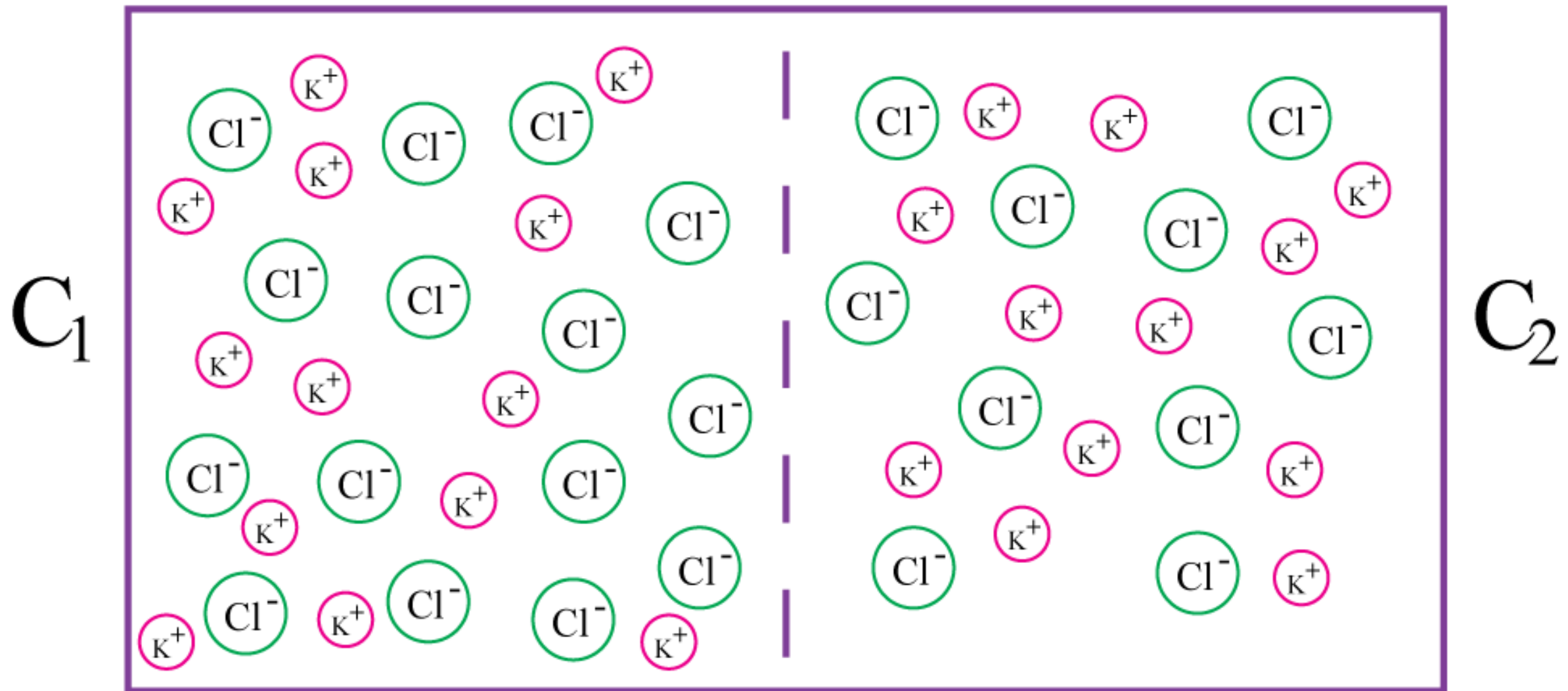


Figure Source: <https://faculty.washington.edu/chudler/ap.html>

Ion concentration difference across membrane lead to potential difference

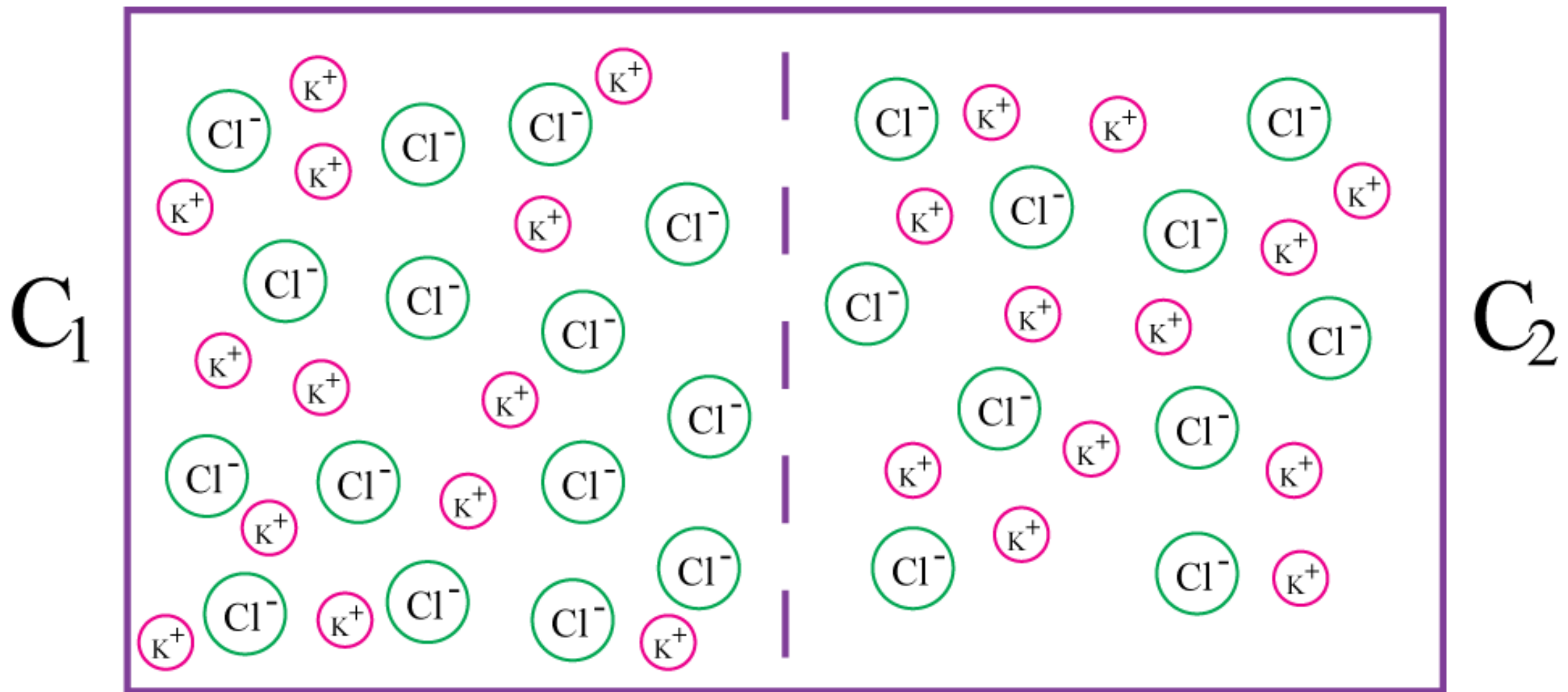
# Nernst Equation

Diffusion of only  $K^+$  ions  
 $\xrightarrow{(C_1 > C_2)}$



# Nernst Equation

Electrostatic attraction on  $K^+$  ions  
←  
( $C_1 > C_2$ )



# Nernst Equation

$$\frac{p_1}{p_2} = \frac{C_1}{C_2} = \frac{e^{-\frac{zeV_1}{k_B T}}}{e^{-\frac{zeV_2}{k_B T}}}$$

$$V_2 - V_1 = \frac{k_B T}{ze} \ln \frac{C_1}{C_2}$$

$z$  is valancy of ion and  $e = 1.6 \times 10^{-19} \text{ C}$

$z = +1$  for  $\text{Na}^+$  and  $z = -1$  for  $\text{Cl}^-$

Ion species	Intracellular concentration (mM)	Extracellular concentration (mM)	Nernst potential (mV)	Important: Nernst Potential is measured w.r.t. potential outside of the cell.
$\text{K}^+$	155	4	-98	
$\text{Na}^+$	12	145	67	
$\text{Ca}^{2+}$	$10^{-4}$	1.5	130	
$\text{Cl}^-$	4	120	-90	

# Summary

- Thermal Energy, Thermal forces and Thermal Motion
- Relative importance of thermal energy
- Boltzmann's law
- Applications of Boltzmann's law