

# BIO-212 - Lecture 14

## Recap of Key Concepts

**Aleksandar Antanasijević, Asst. Prof.**

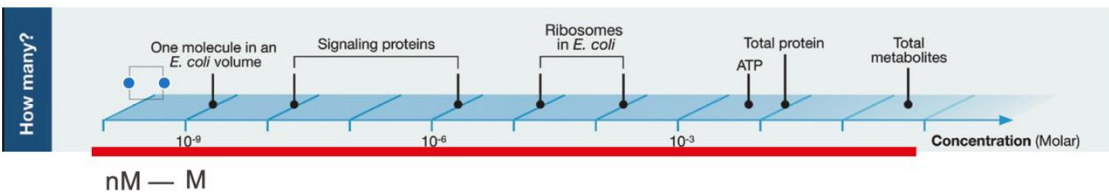
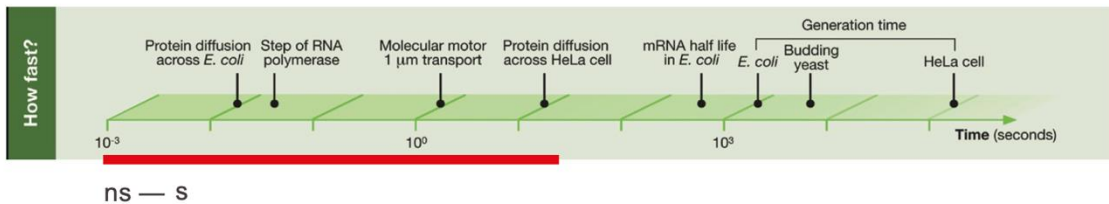
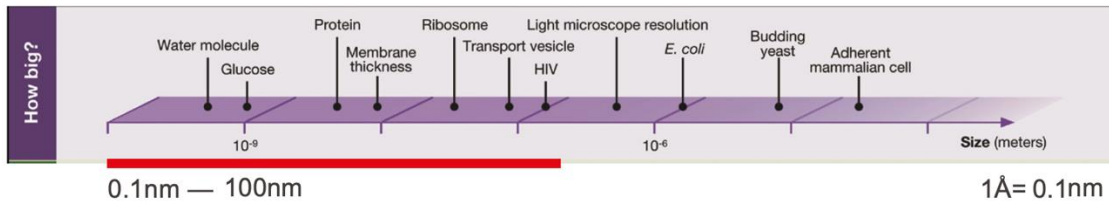
Laboratory of Virology and Structural Immunology  
Global Health Institute, School of Life Sciences  
École Polytechnique Fédérale de Lausanne



17th of December 2025

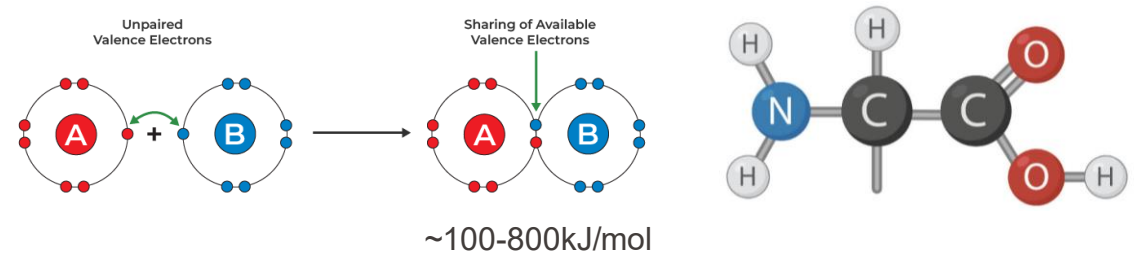
# Lecture 1 – Basic concepts: Atoms and Interactions

• Biomolecules on the scales of life:



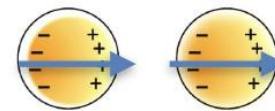
• Atomic and molecular interactions in biomolecules

## Covalent bonds

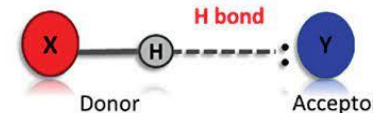


## Non-covalent interactions

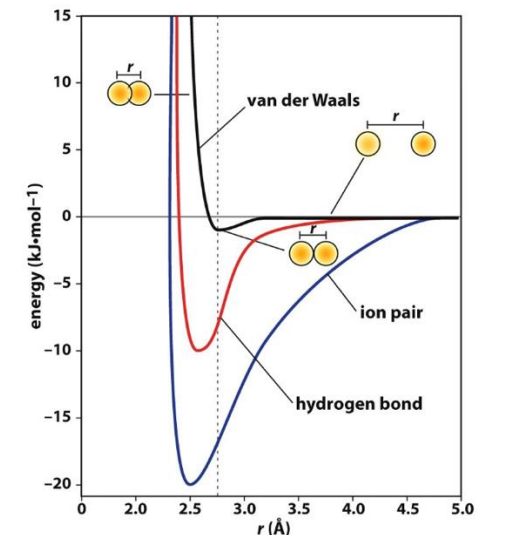
### van der Waals interactions



### Hydrogen bonds



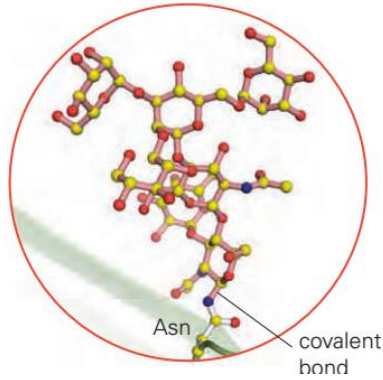
### Ionic interactions



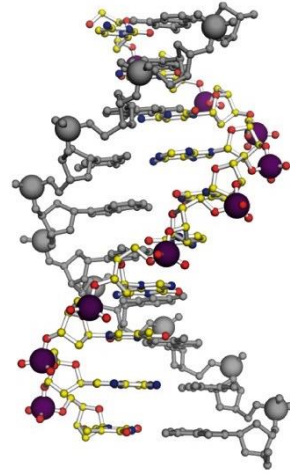
# Lectures 2 - 5: The Molecules of Life

Macromolecular Structure

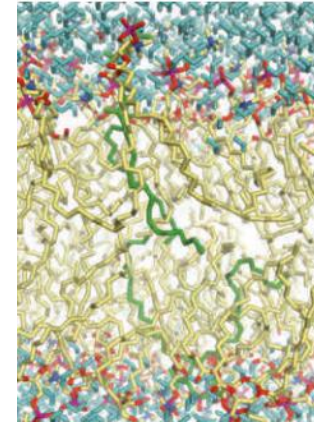
## Carbohydrates



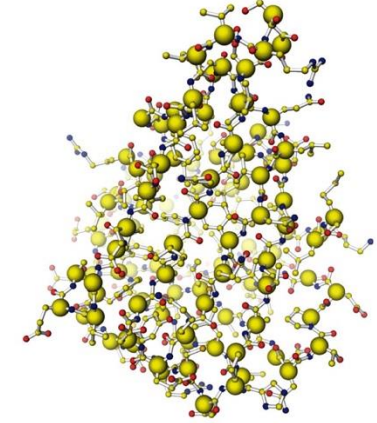
## Nucleic Acids



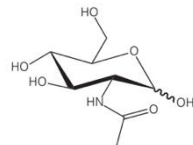
## Lipids



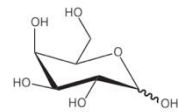
## Proteins



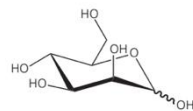
Building Block



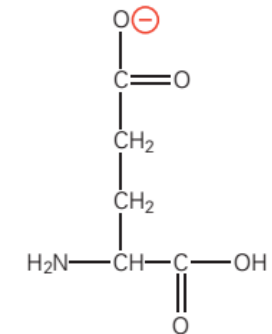
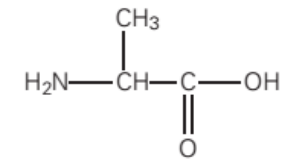
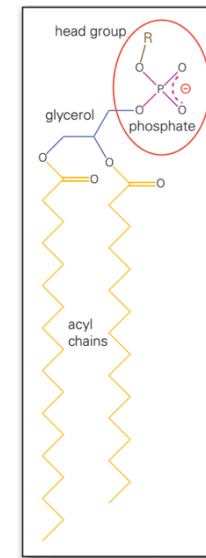
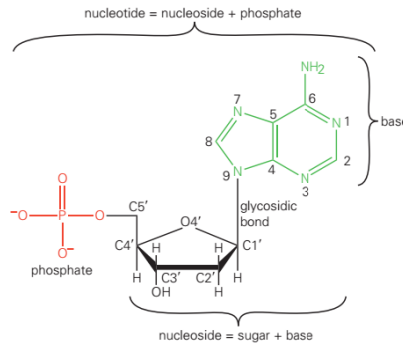
N-acetylglucosamine (GlcNAc)



galactose (Gal)



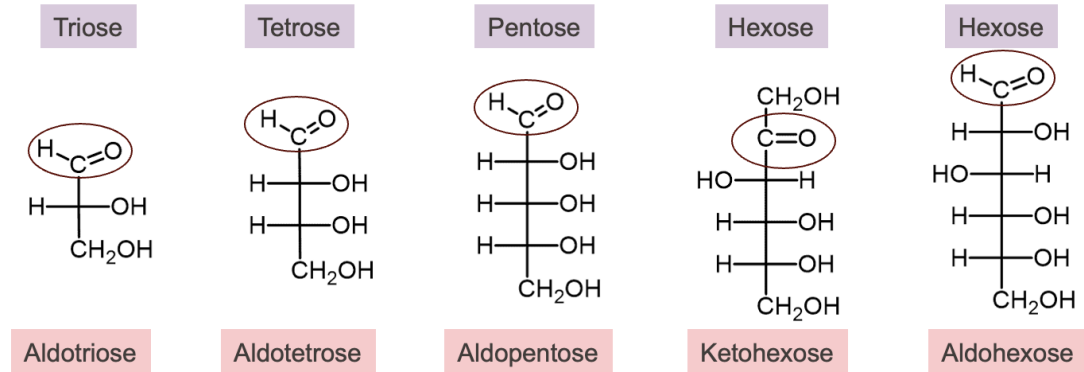
mannose (Man)



# Lecture 2 – Carbohydrates

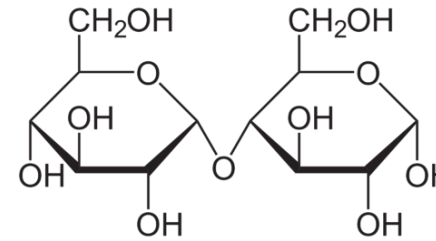
• Carbohydrate building blocks

## Monosaccharides:

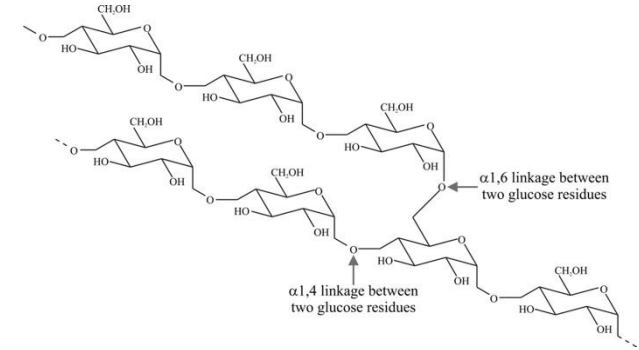


• Carbohydrate polymers and their roles

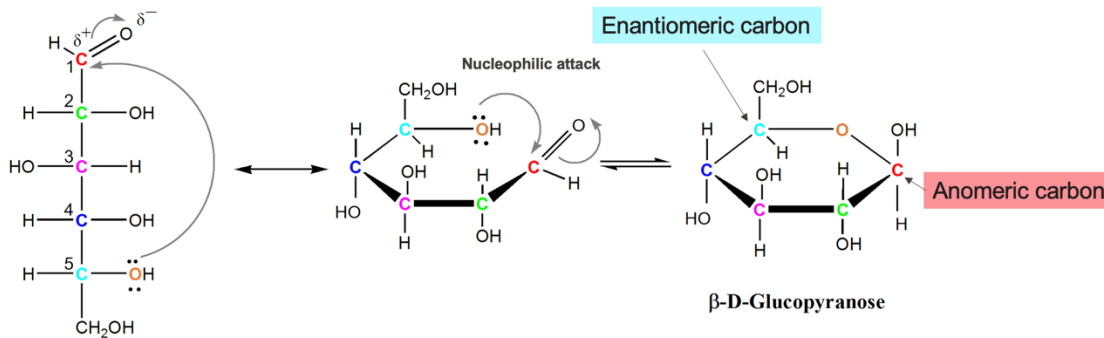
## Disaccharide (Maltose)



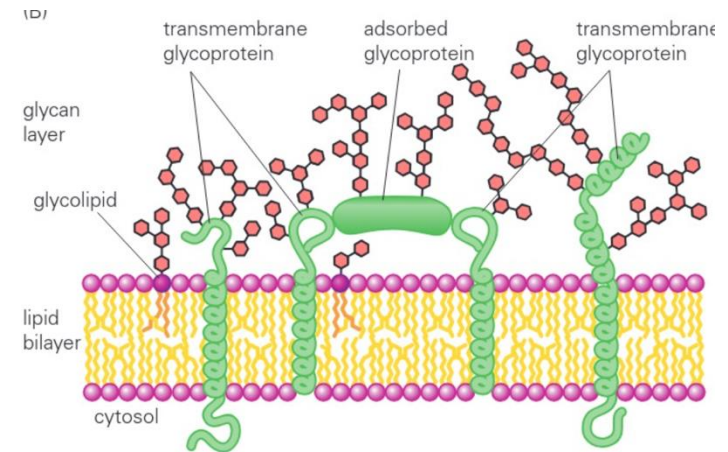
## Polysaccharide (Amylopectin)



## Hemiacetal and hemiketal forms in solution:



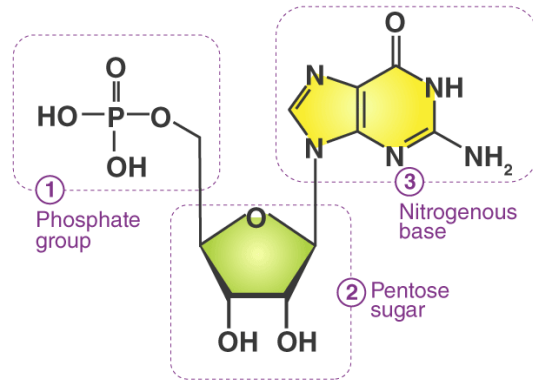
## Glycolipids and glycoproteins



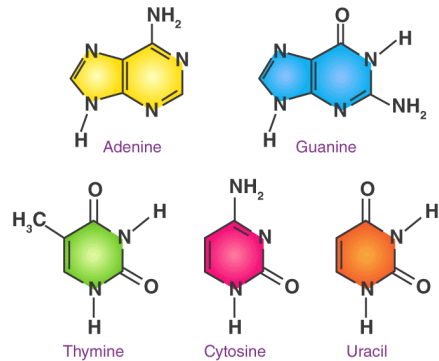
# Lecture 3 – Nucleic Acids

## • Nucleic Acid building blocks

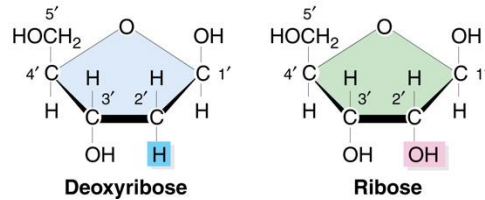
- 3 main components: **base, pentose, and phosphate**



- 5 base options



- 2 pentose options



- 2 x 4 sets of nucleotides to produce DNA and RNA

## • Nucleic Acids (DNA and RNA)

- Linear polymers of nucleotides

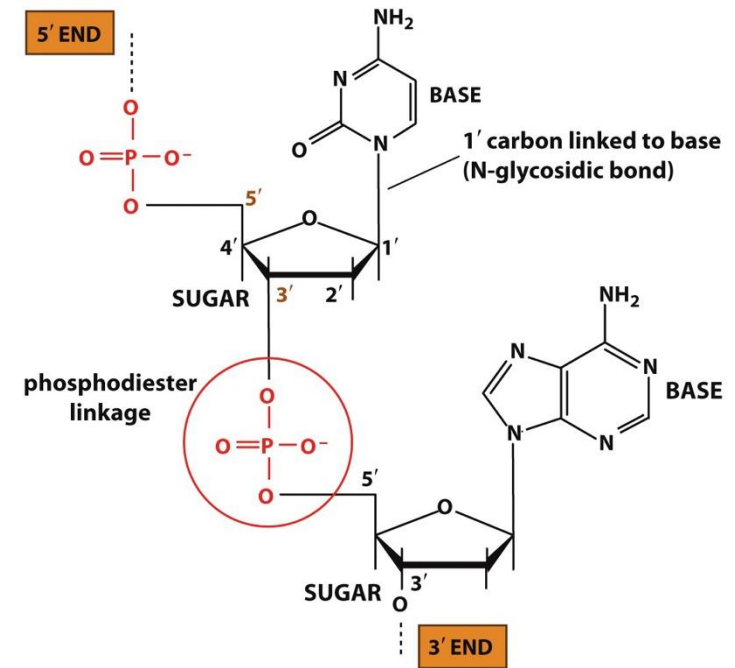


Figure 1.20 The Molecules of Life (© Garland Science 2013)

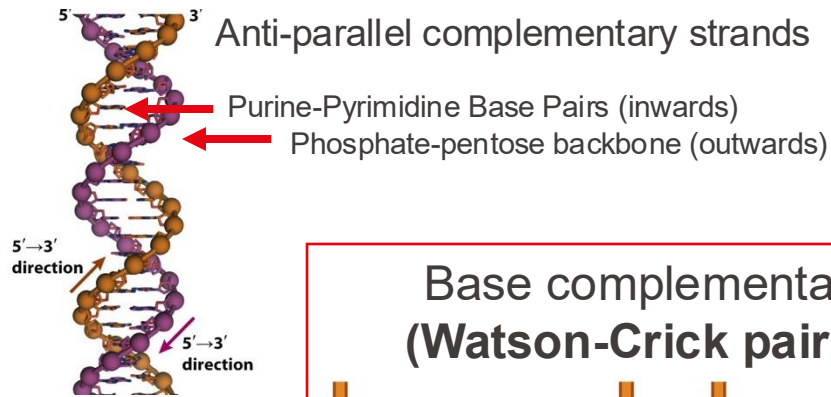
- 5'->3' directionality in addition of nucleotides

- Attachment via phosphodiester linkages

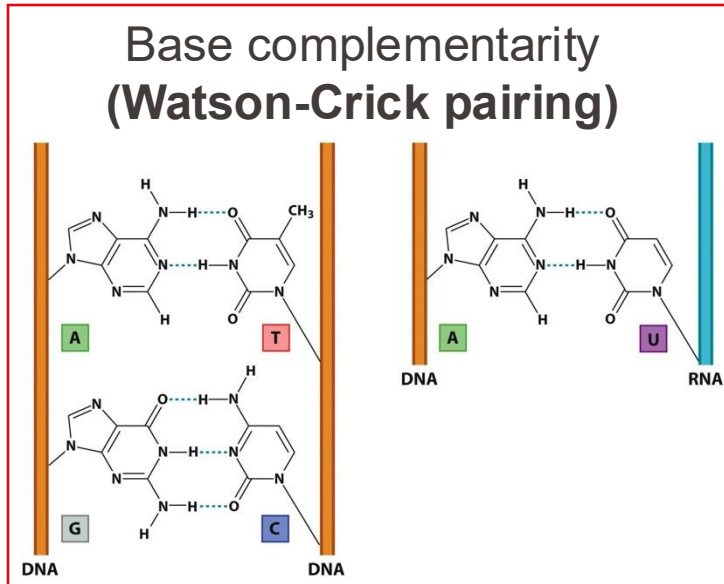
# Lecture 3 – Nucleic Acids

## • Nucleic Acid structure and assembly

- Double-stranded helical assembly of DNA (B form)



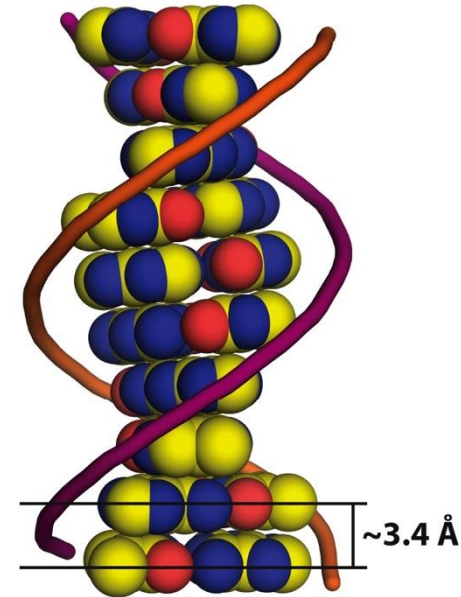
10 base pairs per helical turn



- RNA can be double- and single-stranded and features greater conformational and functional diversity

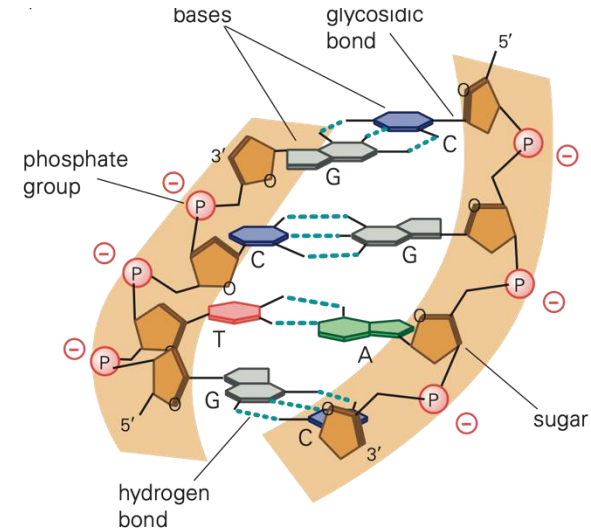
## • Important non-covalent interactions

Van der Waals interactions (layered base stacking)



- Stacking of aromatic  $\pi$ -orbitals ( $\pi$ - $\pi$  stacking)

Hydrogen-bond network (base pairing)

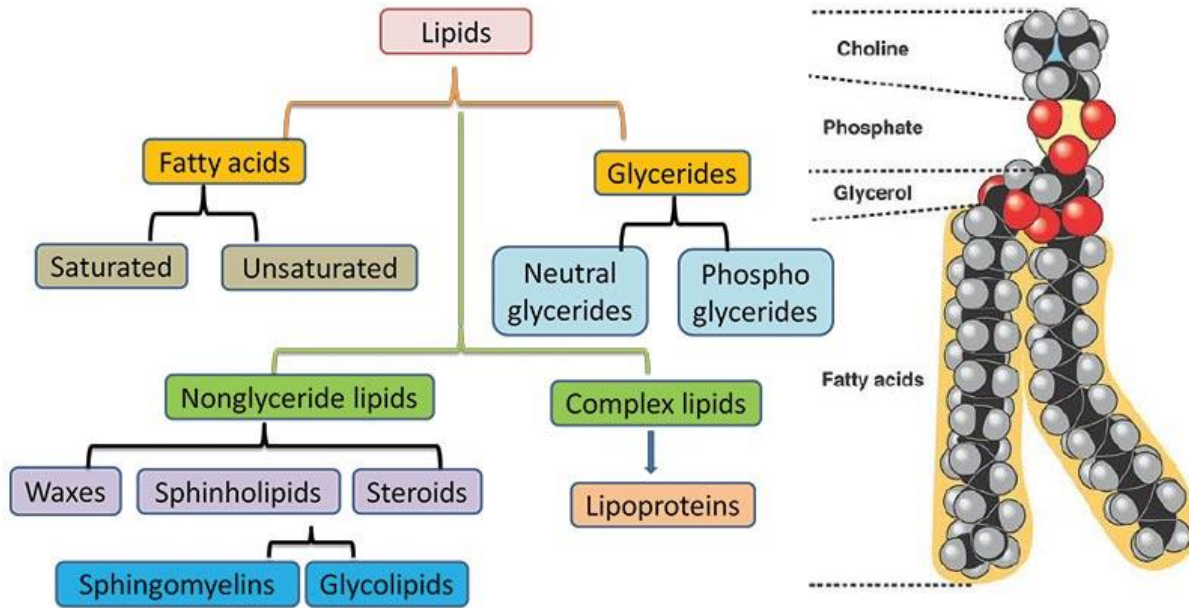


- Charged, polar backbone attracts water and positively charged ions

# Lecture 4 – Lipids

## • Main lipid types and roles

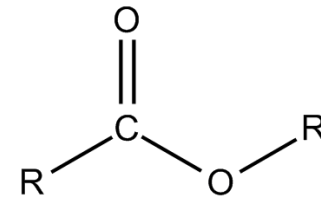
- Main lipid types based on chemical properties



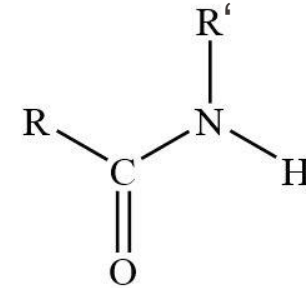
- There are >1000 building blocks which can assemble in different ways (very diverse)
- Their main roles include energy storage, assembly of biological membranes, cell and hormone signaling

## • Important bonds and interactions

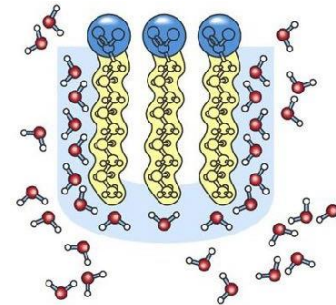
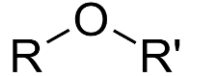
Ester bonds  
(triglycerides)



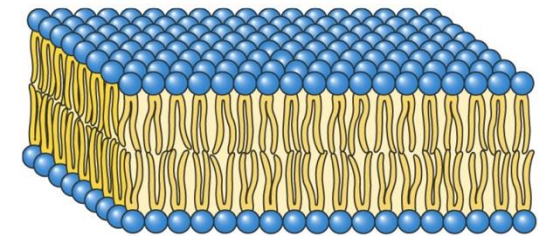
Amide bonds  
(sphingolipids)



Ether bonds  
(head domains)



- Amphipathic molecules (polar and hydrophobic) that form bilayers in aqueous solutions



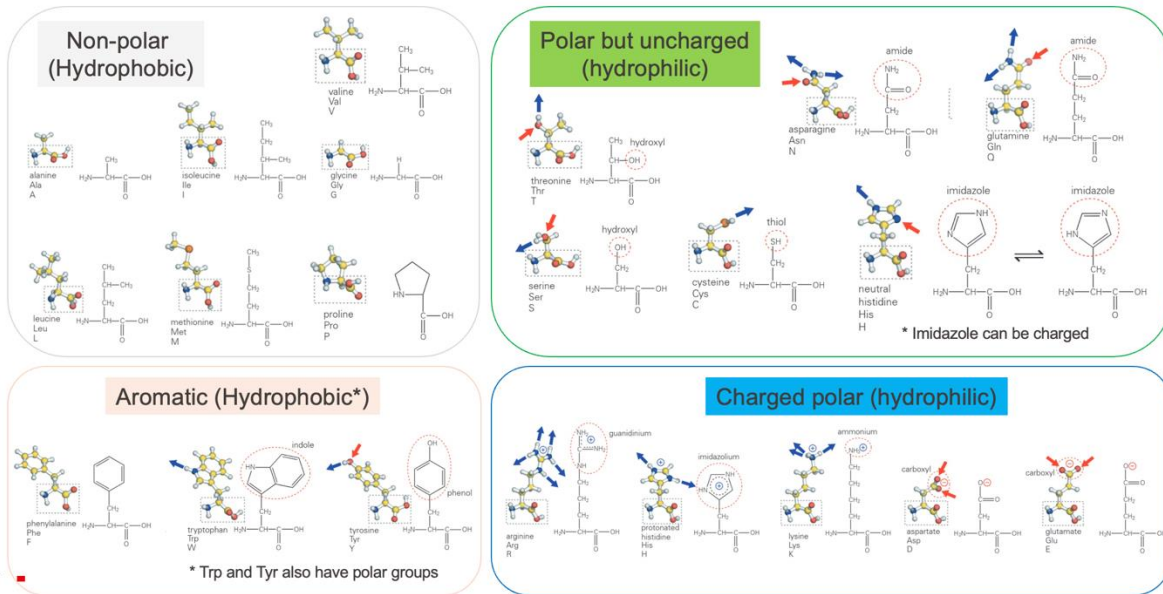
**Bilayer**

- Tail: vdW interactions
- Head: hydrogen bonding and charged interactions

# Lecture 5 – Proteins

## • Physicochemical properties of amino acids

- 20 canonical amino acids that assemble all proteins

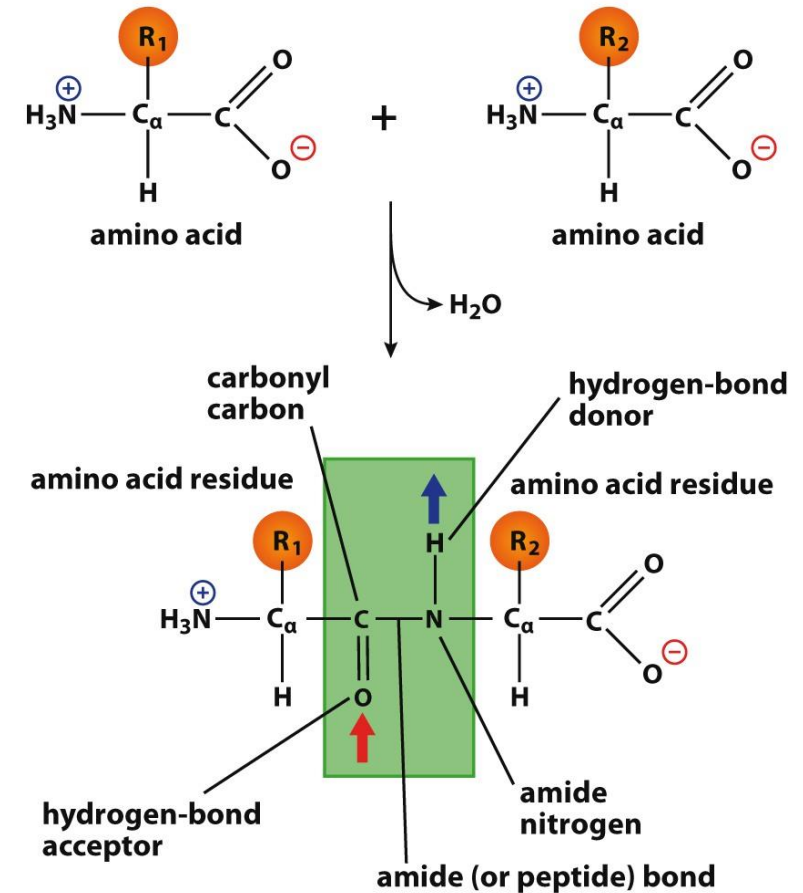


- Their side-chains give them different properties (e.g., water solubility, charge, hydrogen bonding capacity).

- They exist as L- or D- stereoisomers, but the proteinogenic amino acids are all L

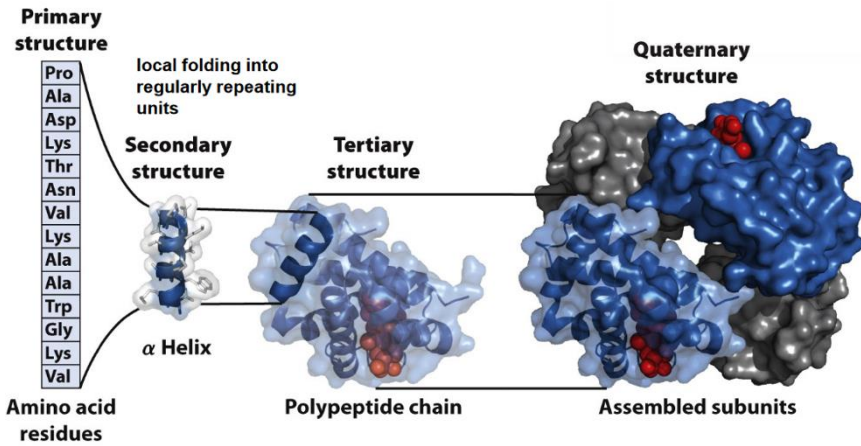
## • Proteins are formed by linking amino acids

- Amino acids are connected via peptide (amide) bonds

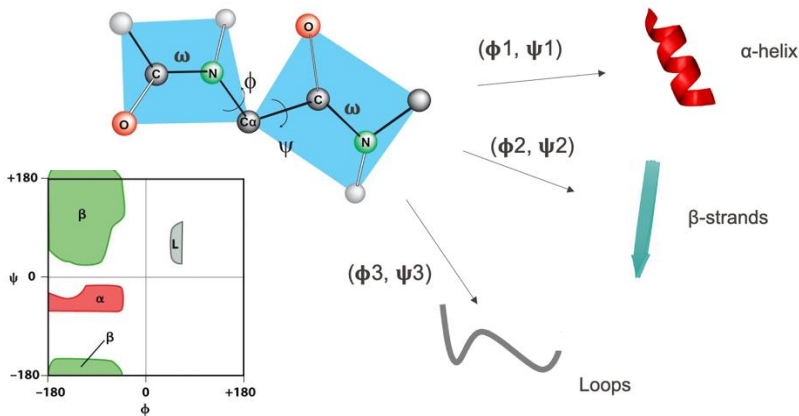


# Lecture 5 – Proteins

- Proteins fold into functional quaternary entities

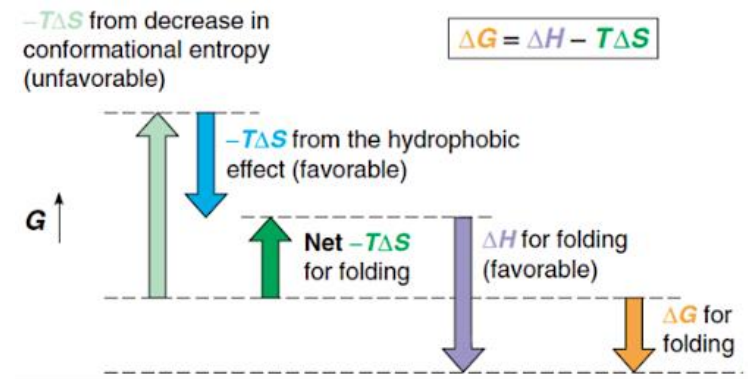


- Torsions around different bonds are predefined by the surrounding chemical groups

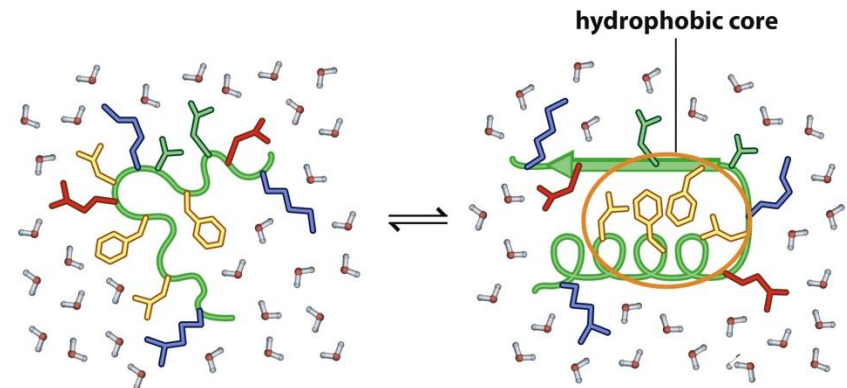


- Protein folding process

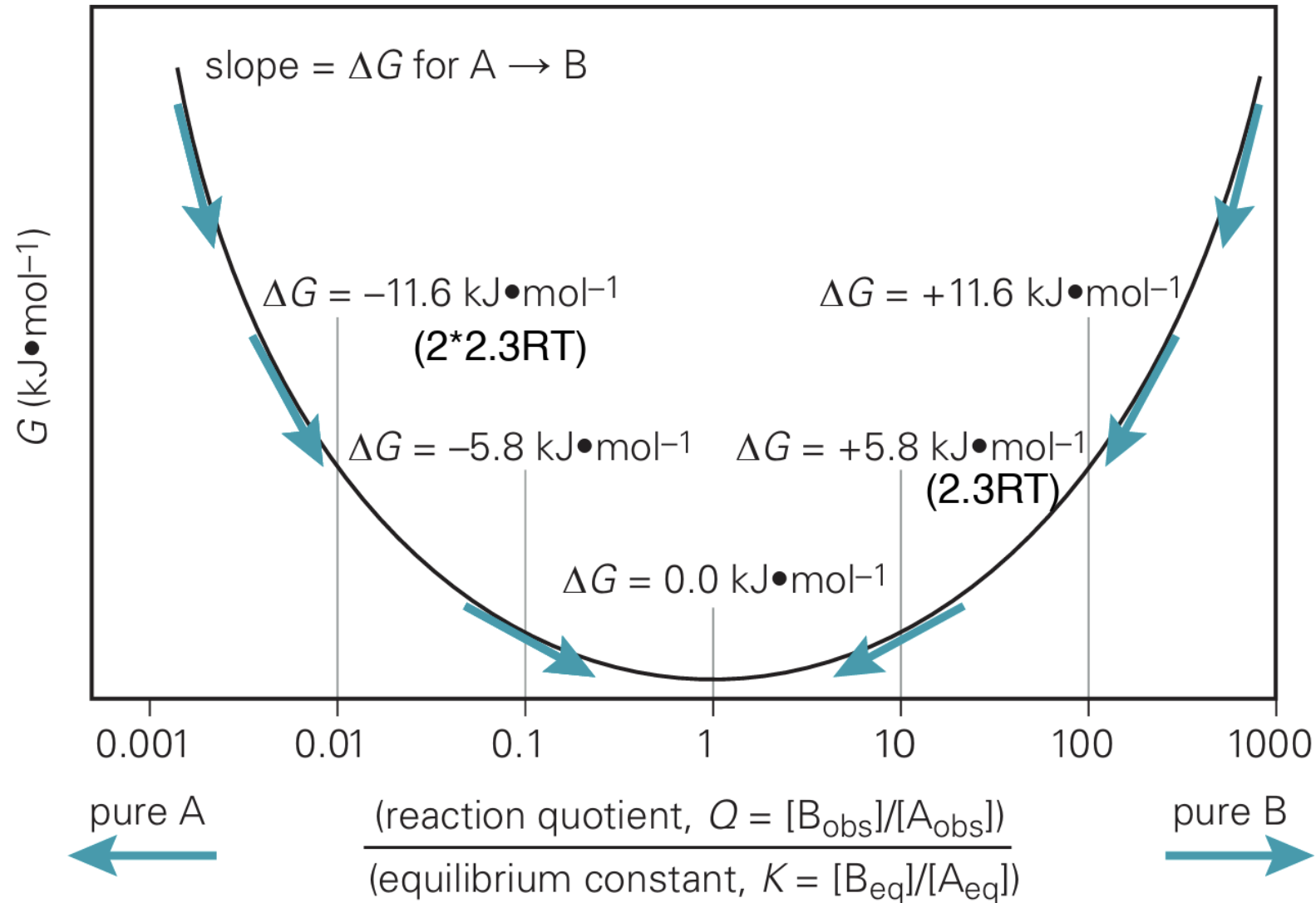
- Protein folding is a process driven by thermodynamic tendency to optimize Gibbs free energy



- Protein folding usually starts with a hydrophobic core



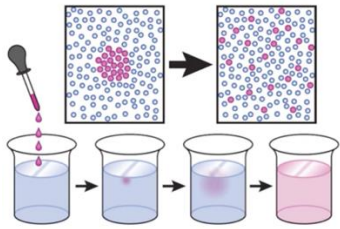
# Lectures 6 - 7: Biomolecule thermodynamics



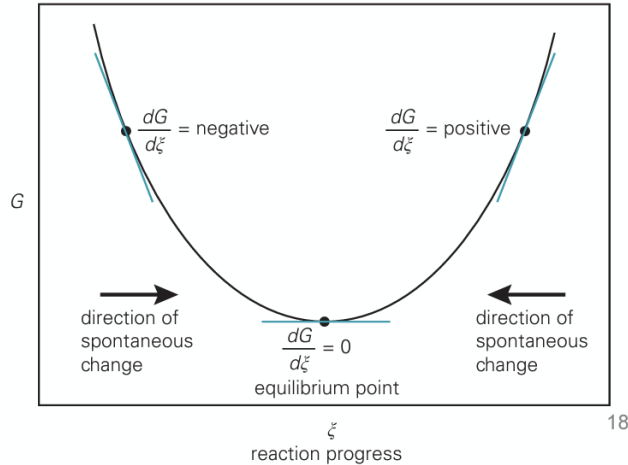
# Lecture 6 – Basic thermodynamic concepts

## Gibbs free energy and chemical potential

- Gibbs free energy ( $\Delta G$ )

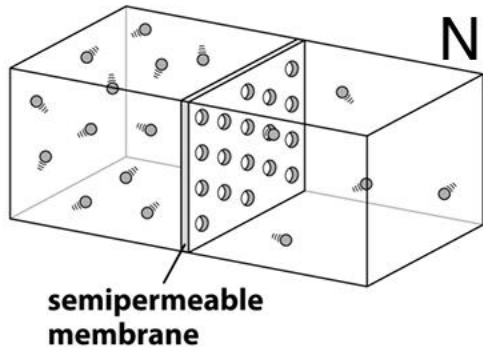


$$dG = dH - TdS$$



18

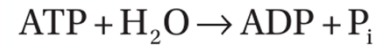
- Chemical potential ( $\mu$ ) defines the concentration conditions for biological processes in equilibrium.



$$\Delta\mu = \mu_2 - \mu_1 = RT \ln \left( \frac{C_2}{C_1} \right)$$

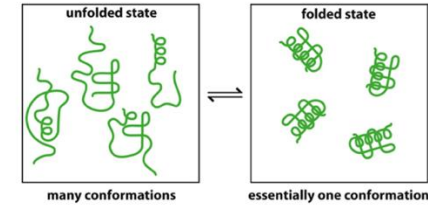
## Equilibrium in different contexts

Chemical reaction



$$K = \frac{[\text{ADP}][\text{P}_i]}{[\text{ATP}]}$$

Protein Folding



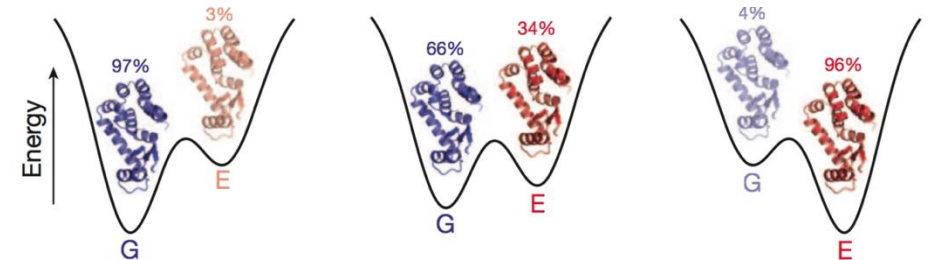
$$K_{\text{folding}} = \frac{[F]}{[U]}$$

Acid-Base Eq.



$$K_a = \frac{[\text{H}^+][\text{A}^-]}{[\text{HA}]}$$

- Boltzmann distribution and equilibrium

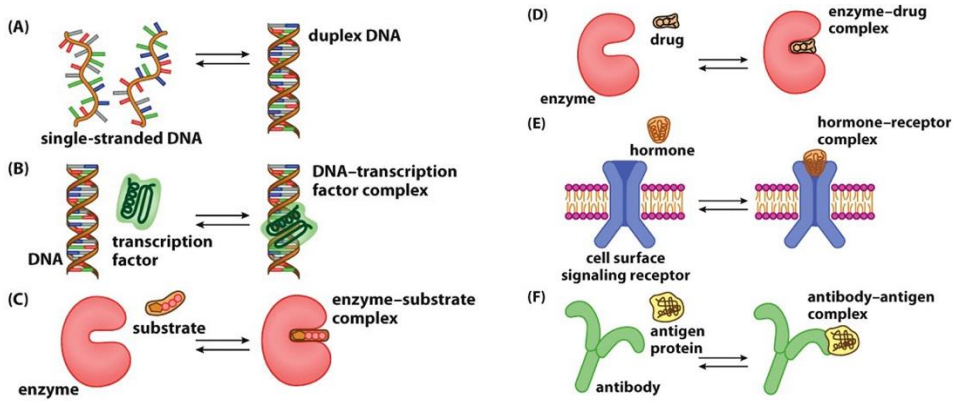


$$\frac{[E]}{[G]} = e^{-\frac{\Delta G_{G \rightarrow E}}{RT}}$$

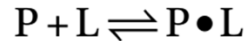
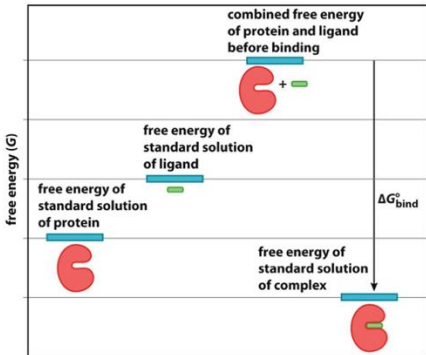
# Lecture 7 – Thermodynamics of molecular interactions

## • Biomolecular interactions (binding)

- Binding in different contexts



- Gibbs free energy and equilibrium constants (K)



$$K_D = \frac{[P][L]}{[P \bullet L]} = \frac{1}{K_A}$$

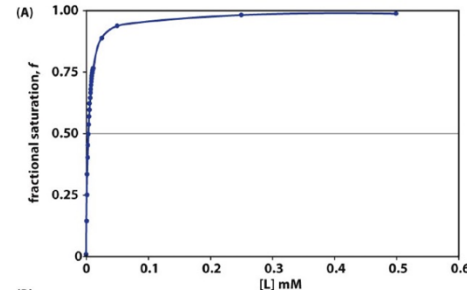
$$\Delta G_{\text{bind}}^0 = -RT \ln K_A$$

$$\Delta G_{\text{bind}}^0 = +RT \ln K_D$$

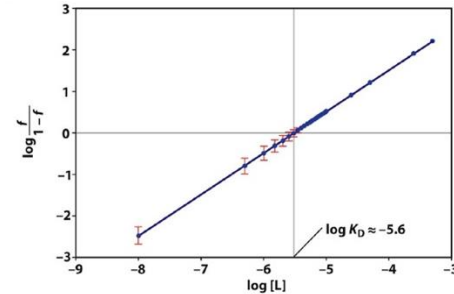
## • Different ways of presenting binding curves

- Fractional saturation (f):

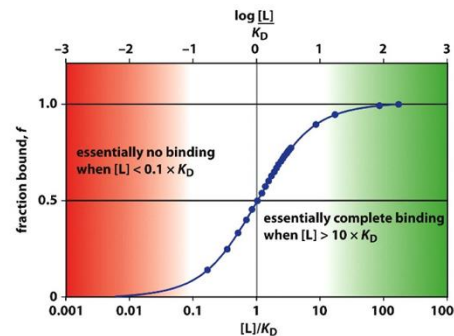
$$f = \frac{\text{concentration of protein with ligand bound}}{\text{total protein concentration}}$$



$$f \sim [L]$$

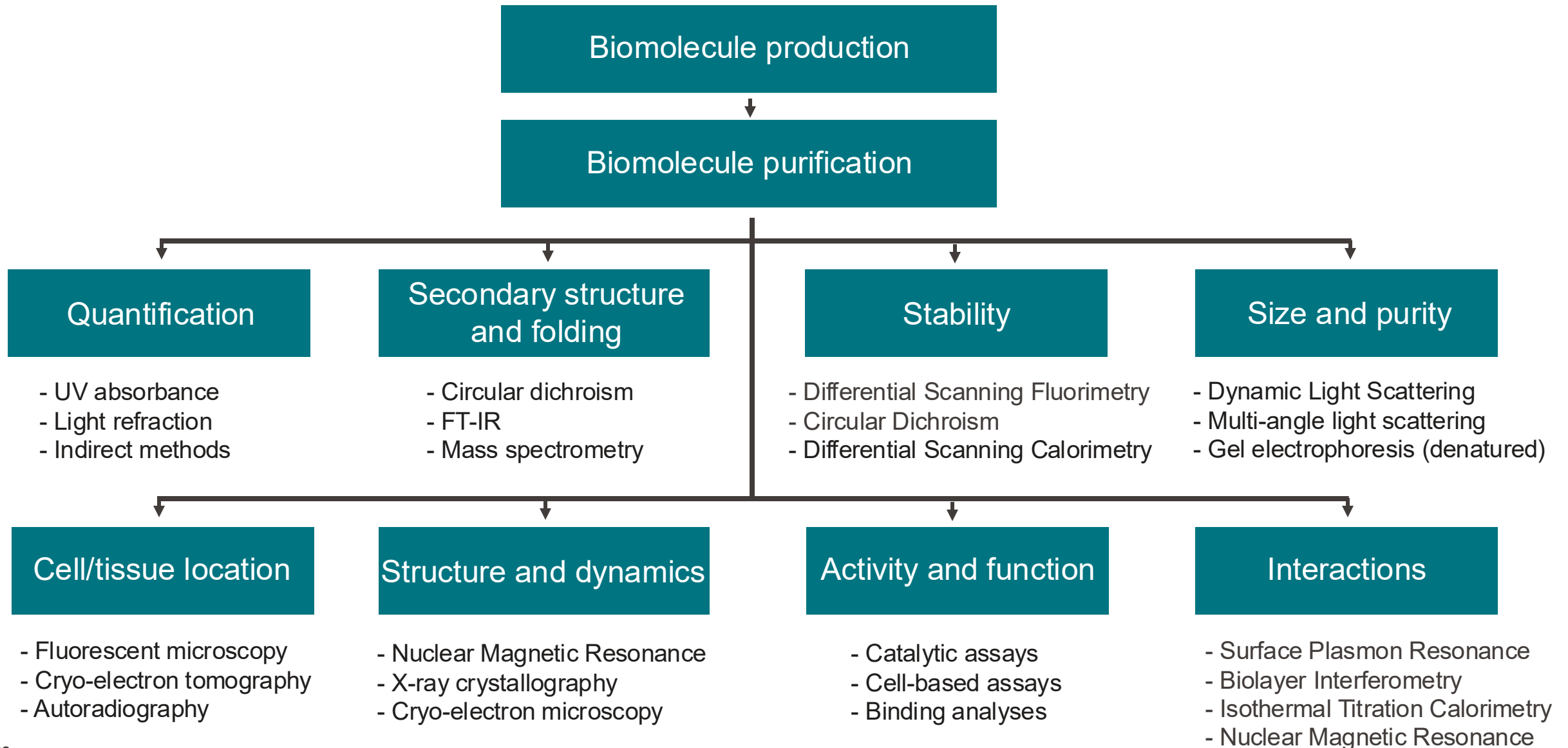


$$\log \frac{f}{1-f} \sim \log [L]$$



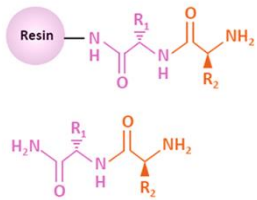
$$f \sim [L]/K_d$$

# Lectures 8 - 13: Working with biomolecules

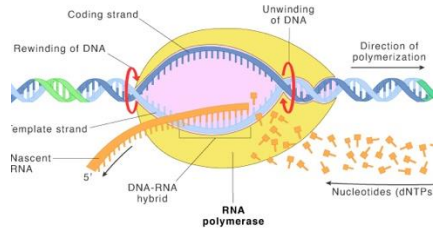


# Lecture 8 – Biomolecule Production and Purification

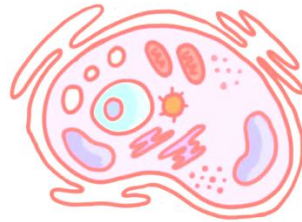
• Biomolecule production methods



Chemical

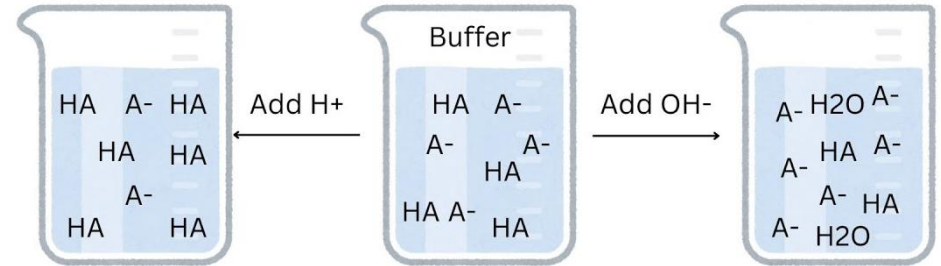


Enzymatic



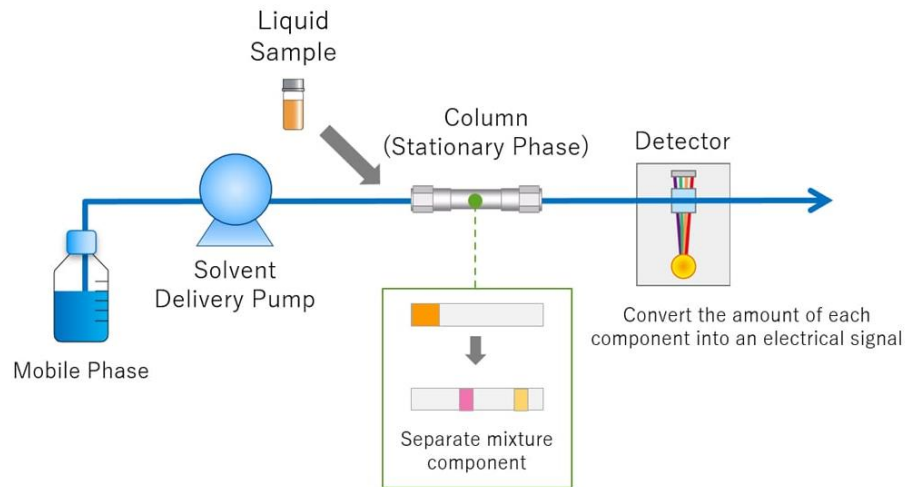
Cell-based

• Buffers and buffer components

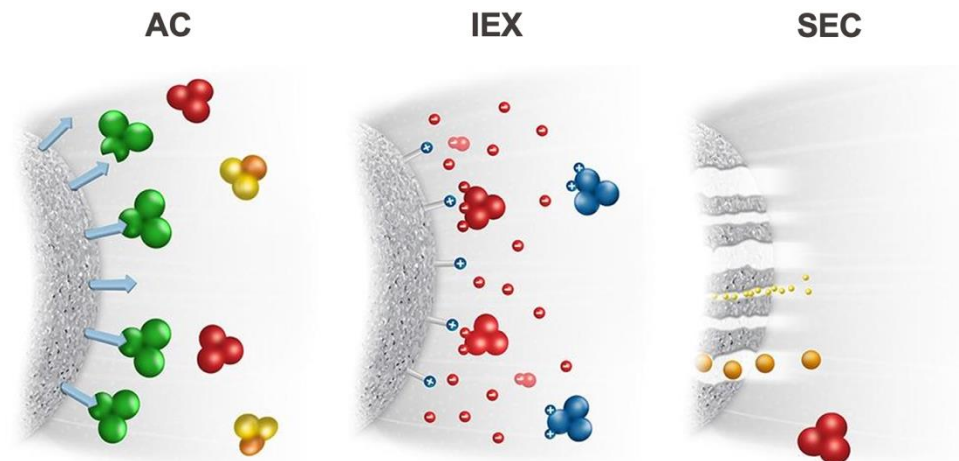


- Maintaining pH and ionic strength of the solution
- Other components can be added for LC or stability

• Liquid Chromatography for biomolecule purification

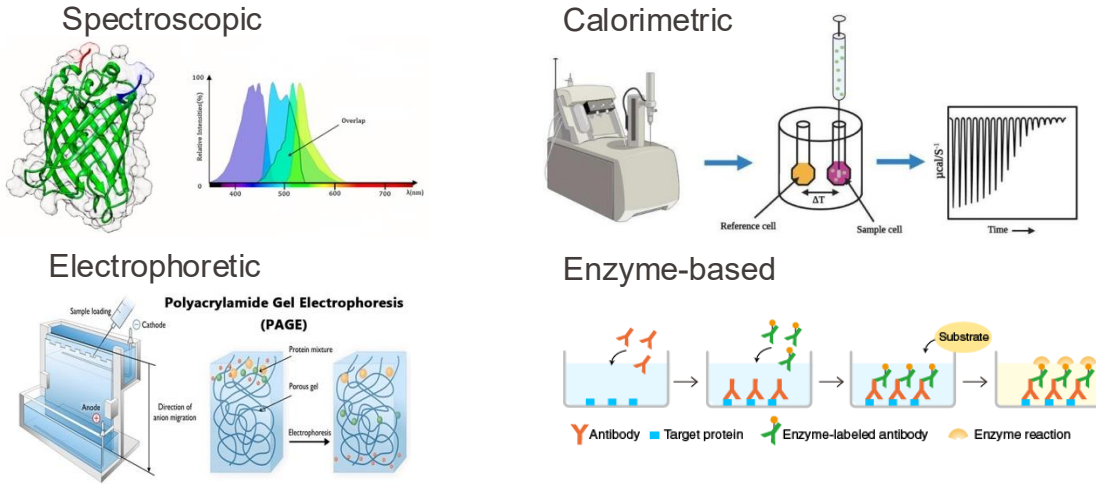


• Basic chromatography methods

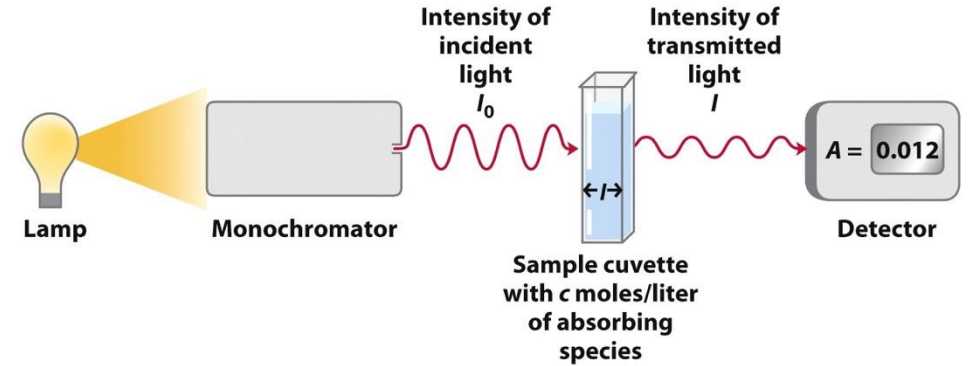


# Lecture 9 – Biomolecule Characterization

## • Different types of measurements

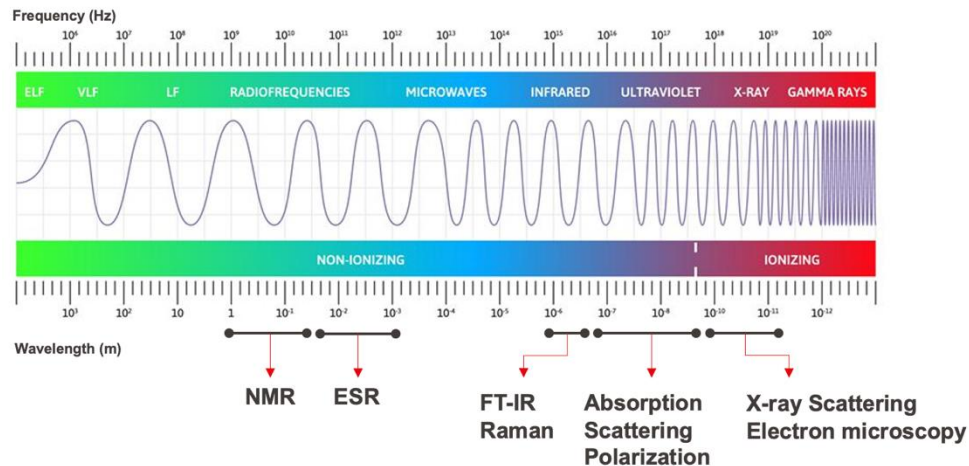


## • Spectroscopic / Spectrometric methods

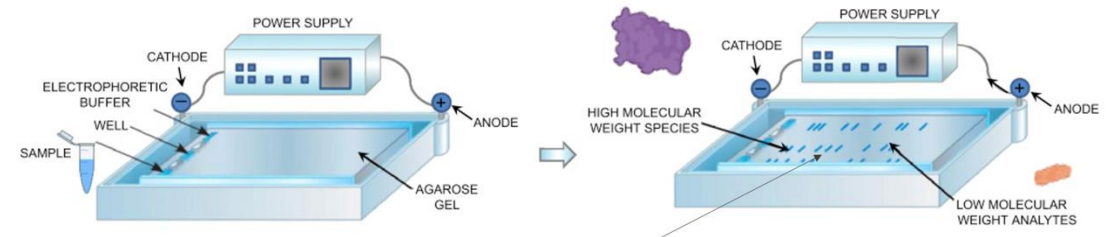


UV/Vis, FT-IR, CD, Fluorescence, DSF, DLS

## • EM spectrum and biophysical methods



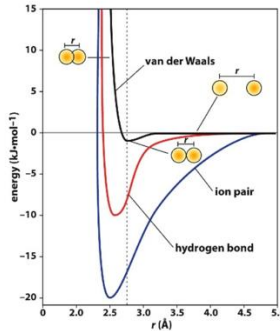
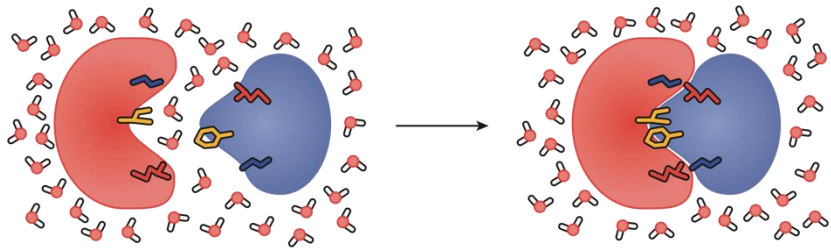
## • Gel electrophoresis



SDS PAGE, Agarose gels

# Lecture 10 – Measuring Biomolecular Interactions

## • Biomolecular interactions and binding



- Small energy contributions from hydrogen bonds, vdW, ionic and hydrophobic interactions
- Water can have a positive and negative impact on binding

## • Dissociation constants and affinity

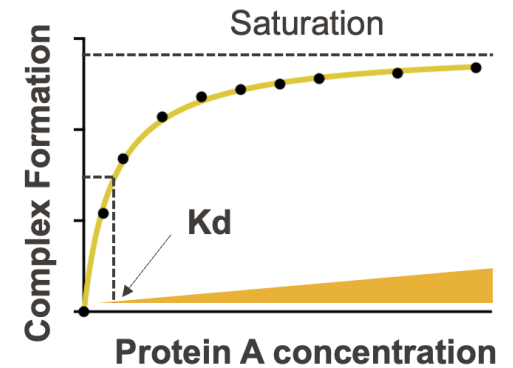
$$\Delta G^\circ = RT \ln K_D = \Delta H - T\Delta S$$

$$\frac{k_d}{k_a} = \frac{k_{off}}{k_{on}} = \frac{[P][L]}{[P \cdot L]} = K_D = \frac{1}{K_A}$$

- Thermodynamic and kinetic analysis of binding

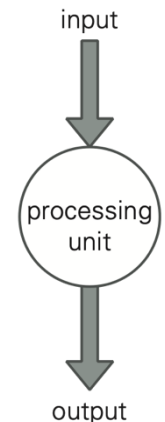
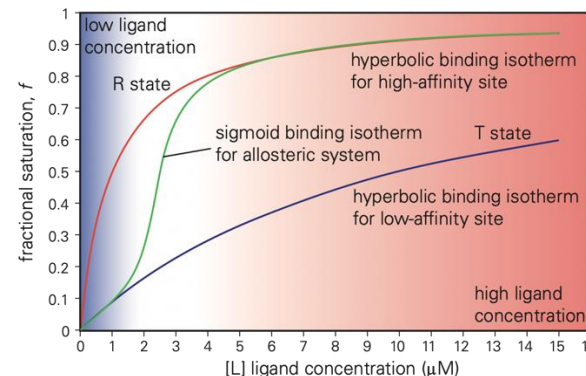
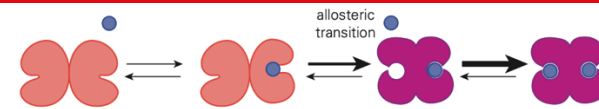
## • Experimental methods for Kd determination

- Titrate one binding partner and measure complex formation



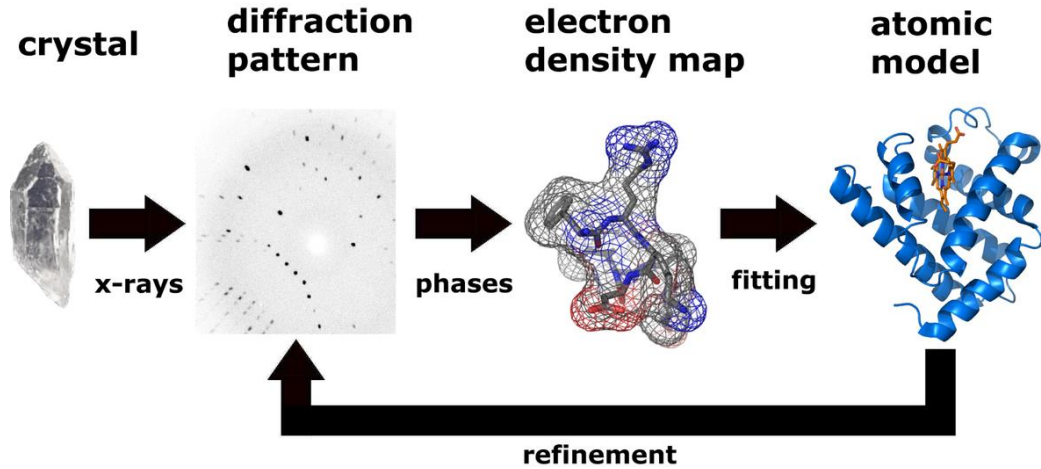
- Calorimetric (ITC) or spectroscopic (FP, SPR, NMR) measurements

## • Cooperativity and Allostery

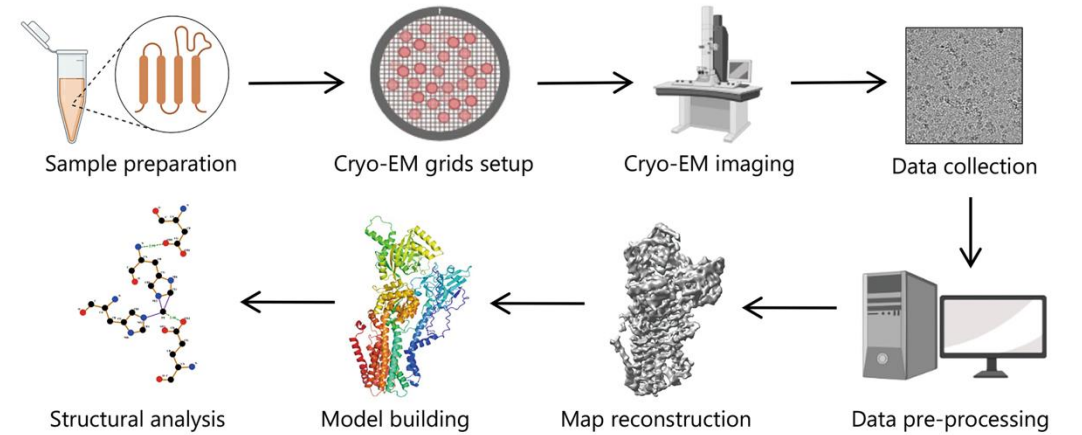


# Lecture 11 – Structural Biology

## • X-ray crystallography



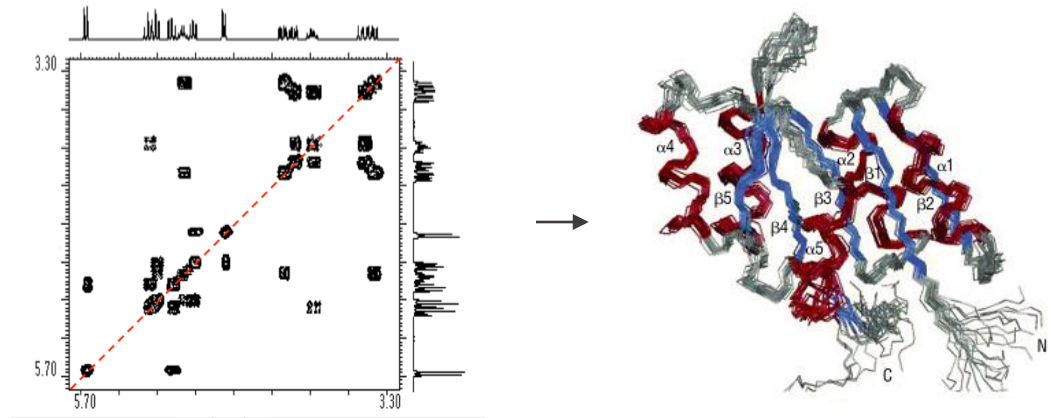
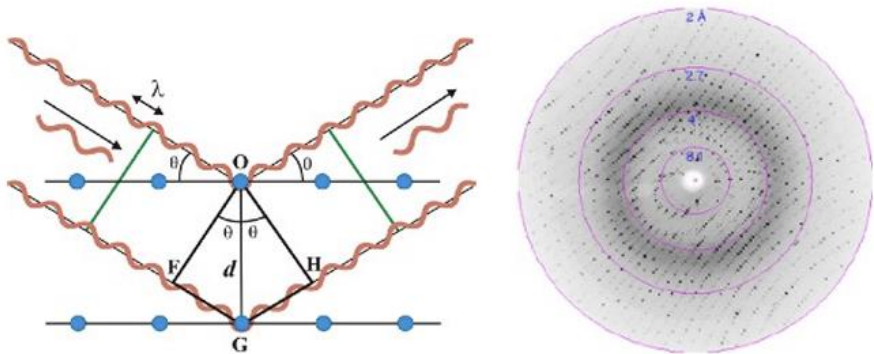
## • Cryo-electron microscopy (CryoEM)



## • Nuclear Magnetic Resonance (NMR)

Bragg's law

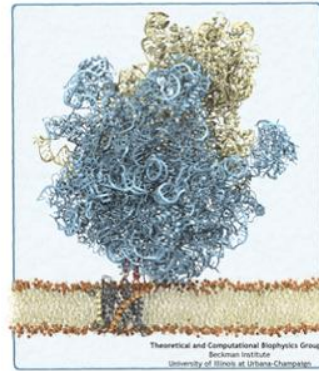
$$n\lambda = 2d \sin\theta$$



# Lecture 12 – Computational Structural Biology

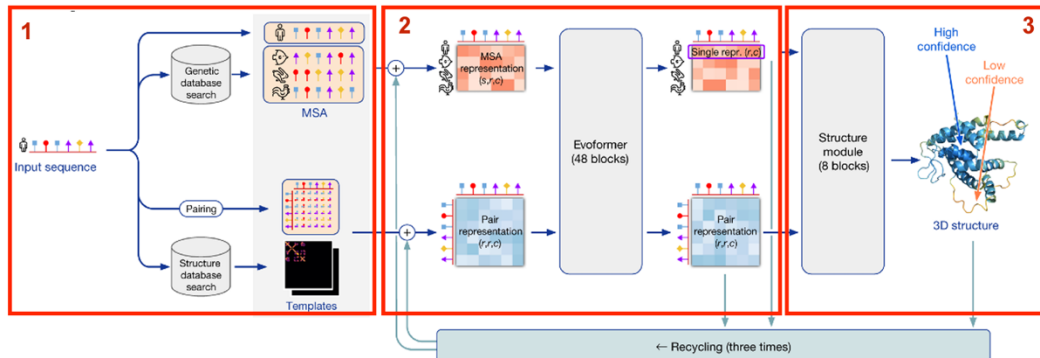
## • Molecular simulations

$$U = \sum_{\text{All Bonds}} \frac{1}{2} K_b (b - b_0)^2 + \sum_{\text{All Angles}} \frac{1}{2} K_\theta (\theta - \theta_0)^2 + \sum_{\text{All Torsion Angles}} K_\phi [1 - \cos(n\phi + \delta)] + \sum_{\text{All nonbonded pairs}} \epsilon \left[ \left( \frac{r_0}{r} \right)^{12} - 2 \left( \frac{r_0}{r} \right)^6 \right] + \sum_{\text{All partial charges}} \frac{332 q_i q_j}{r}$$



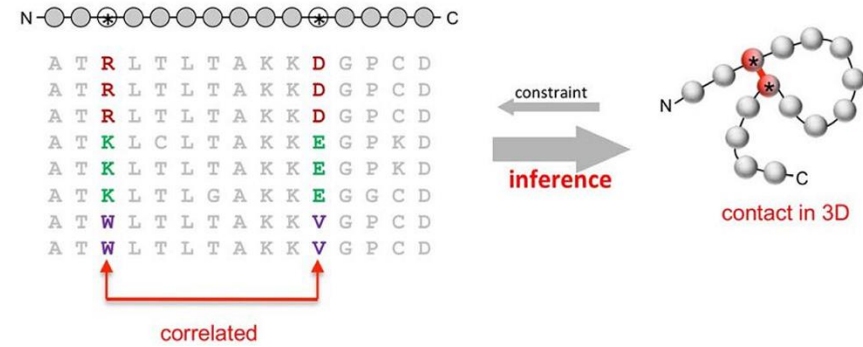
- Modeling biomolecular behavior using physical potentials

## • Structure prediction using AlphaFold 2



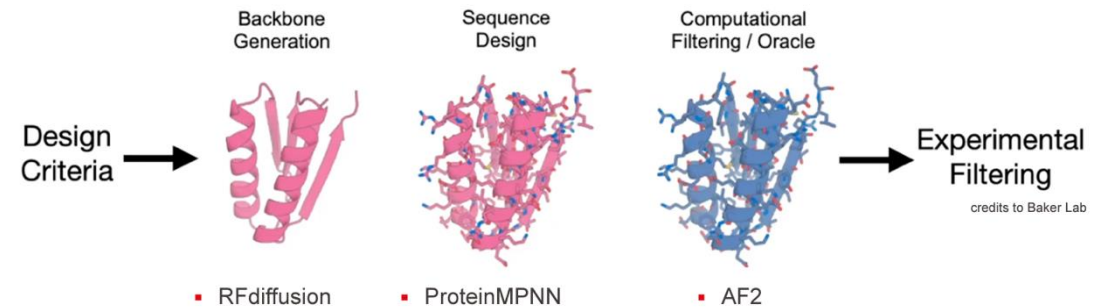
- Machine learning for predicting biomolecule assembly

## • Evolutionary couplings



- Extracting structural information from sequence data

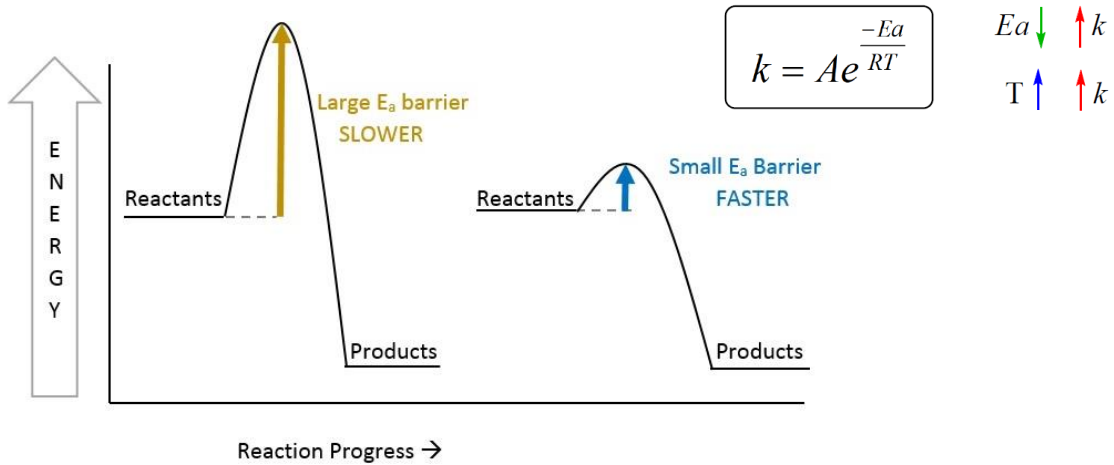
## • Protein design



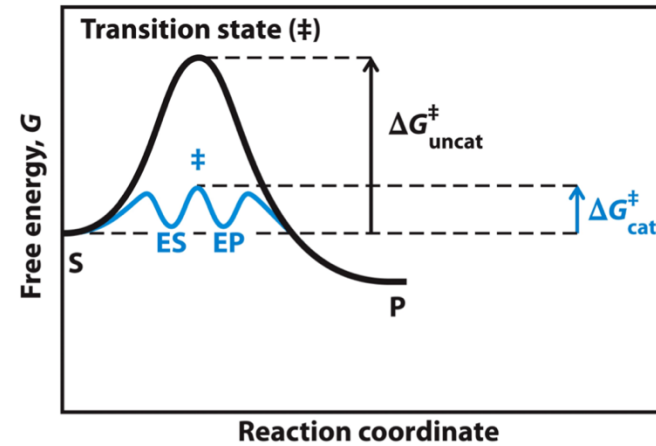
- Machine-learning-based methods for molecule engineering

# Lecture 13 - Kinetics and Catalysis

## • Reaction rates and Activation Energies



## • Catalysis and enzymes

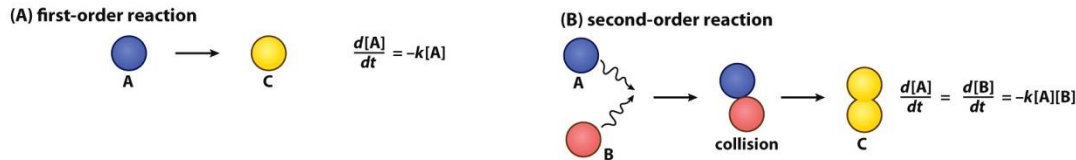


$$V_0 = \frac{V_{\max} [S]}{[S] + K_m}$$

- Michaelis-Menten equation

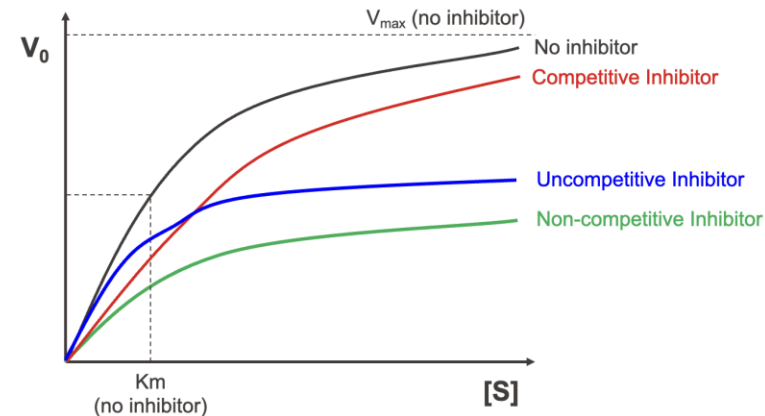
## • Reaction order

- Order of the reaction depends on the number of molecules whose quantities impact the reaction rate



- Rate constant units depend on the order

## • Enzyme inhibition mechanisms



- Inhibition depends on the  $K_i$  and inhibitor conc.

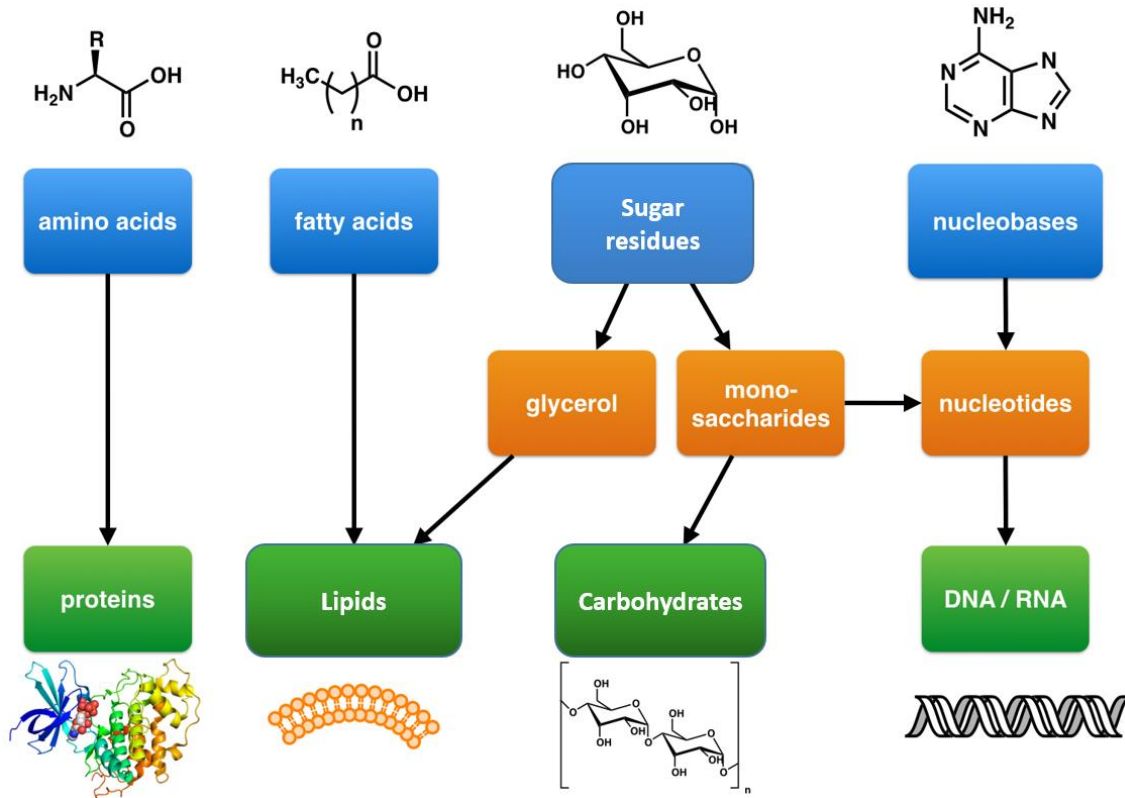
- Often expressed as  $\alpha$

$$\alpha = 1 + \frac{[I]}{K_i}$$

# Essential content

- The **theory** from all lectures (fundamental concepts, mechanisms, properties of biomolecules, correlations...).
- Biomolecule **assembly** and their **properties** (chemical bonds and their properties, relevant functional groups, non-covalent interactions...)
- Biomolecule **production** and **purification** (principles, systems for production, chromatographic methods...)
- **Thermodynamic** principles, **biophysical** methods and **computational** approaches applied in biochemistry
- **Equations** for calculating constants, energies, reaction rates etc.
- **Exercises** (questions, problems, calculations...)

# How to approach learning about biomolecules?



What are the building blocks?

What are the most important covalent bonds?

What are the most important non-covalent interactions?

How do they assemble in 3D space?

What function do they serve in cells?

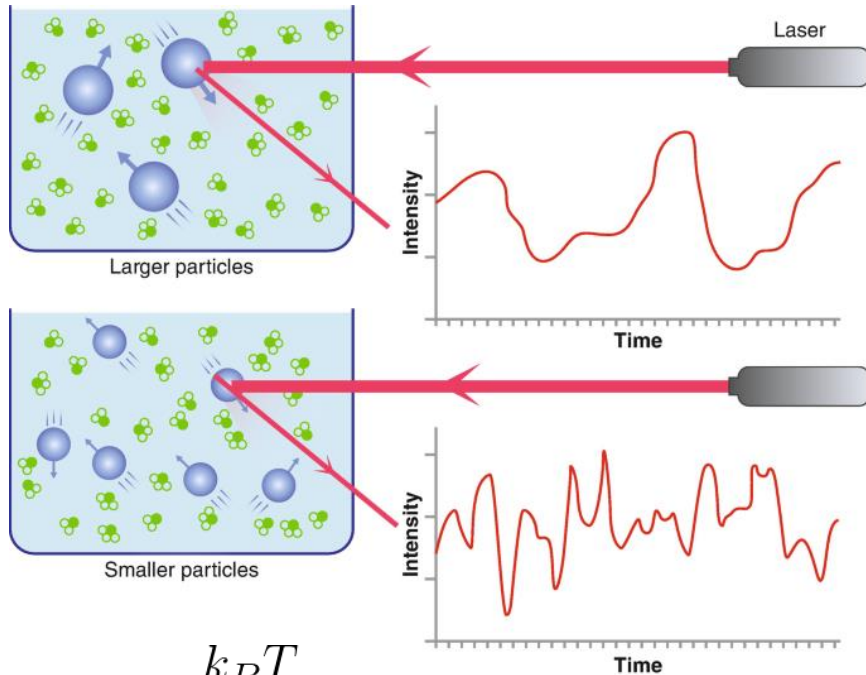
What are some important examples?

Are there subtypes? How do they differ?

Molecular structures of the essential building blocks of **proteins**, **carbohydrates** and **nucleic acids** need to be learned!

# How to approach learning about experimental methods?

## Example: Dynamic Light Scattering



$$R_H = \frac{k_B T}{6\pi\eta D}$$

How do different experimental methods work?

What are the quantities they measure?

What are their target biomolecule applications?

What is the minimal requirement for the method to work?

What are the limitations in their applicability?

Are there alternatives that cover the gap in applicability?

All relevant equations used for calculations of molecular properties need to be learned for the exam!

# Exam Details

## When?

**January 15<sup>th</sup>, 2026**  
**Start time: 15:15**

\* Please be there by 14:45 PM

## Where?

**SwissTech Conv. Center**  
**Main hall: Zones I, J, and M**

Seating will be assigned

## How long?

**3 hours total**  
**End time: 18:15**

Students that requested special accommodations  
will have 1 extra hour (End time: 19:15)

## What to bring?

- 1) Camipro card (VERY IMPORTANT)**
- 2) Writing supplies (pens, pencils etc.)**
- 3) Simple calculator (no smartphone)**

All paper supplies will be provided by the staff

## Exam contents

- 1) Multiple choice questions**
- 2) Short questions/problems**
- 3) Longer multistep problems**

The exam will be in French and English and you can answer in any language

**Books, printouts, cheat-sheets, calculators with memory and fitting capability are not allowed**

