

Enzymes

→ **Proteins** that catalyze, accelerate, speed up, increase the rate of chemical reactions
Small amount
Unconsumed, unchanged
So, they can be used repeatedly

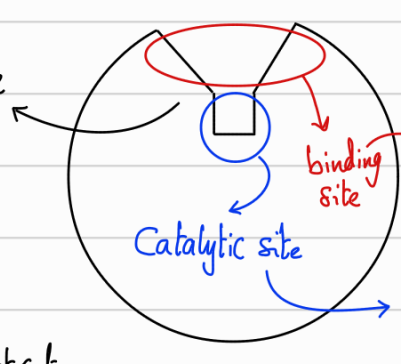
→ Except ribozymes

Affinity → strength of binding
Specificity → The preference to bind a molecule
most proteins are specific
Albumin is non-specific

Active site

Active site must be specific with high affinity, by:

- 1) Compatible 3D shape
- 2) proper interactions with substrate



bind, orient and stabilize the interactions with substrates (reactants)
Catalyze, carry out the reaction

determined by the primary structure (amino acid sequence) of the enzyme

- Active site
- small
 - must bind the substrate on 3 different points to recognize chirality (isomers)
 - Internal, and exclude water
 - Interactions are non-covalent (weak) but multiple (high affinity)

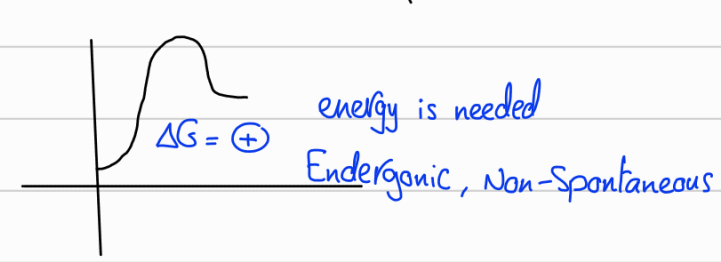
Binding between substrate and active site:

- 1) lock and key model → fits directly
 - 2) Induced fit → induce conformational changes
- Glucokinase bind Glucose (lock and key) and ATP (induced fit)

Free energy

potential energy stored in bonds

Free energy change (ΔG) is energy available for the reaction ($\Delta G = G_{\text{Products}} - G_{\text{Reactants}}$)



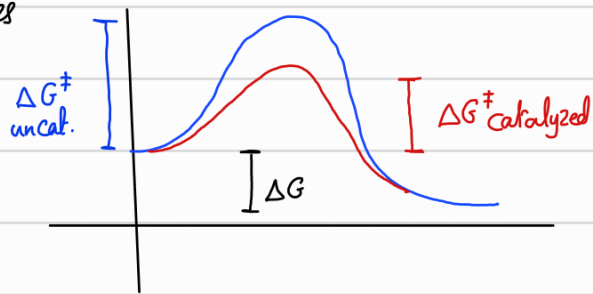
☆ When $\Delta G = \text{zero}$ → Equilibrium (Rate forward reaction = Rate Backward reaction)

☆ Any reversible reaction ($\Delta G_{\text{forward}} = \Delta G_{\text{backward}}$) and opposite in charge

Transition state → intermediate between substrates and products with highest energy
(ES) unstable with maximum strained bonds

Activation energy (ΔG^\ddagger): $G_{\text{Transition state}} - G_{\text{substrates}}$

Enzymes decrease activation energy
but doesn't affect free energy change (ΔG)



Mechanisms of catalysis:

- 1) **Bond strain** \rightsquigarrow bonds become weaker (vulnerable) and broken easier, such as lysozymes
- 2) **Acid / base catalysis** \rightsquigarrow involve proton transfer, such as serine proteases (trypsin, chymotrypsin)
- 3) **Covalent catalysis** \rightsquigarrow formation of covalent bond such as serine proteases (Elastase)

Types of interactions with active site $\left\{ \begin{array}{l} \text{binding site} \rightarrow \text{only non covalent} \\ \text{catalytic site} \rightarrow \text{Covalent or non covalent} \end{array} \right.$

Enzymes Classifications

1) Oxidoreductase \rightsquigarrow Oxidation Reduction Reaction

Dehydrogenase ☆ transfer electrons in the form of H^- or H atom ☆ Require NAD^+ , FAD , FMN ☆ Example: Glutathione reductase	Oxidase ☆ transfer H into O_2 ☆ produce H_2O_2 Hydrogen peroxide \hookrightarrow	Oxygenase ☆ introduce Oxygen Monooxygenase \rightsquigarrow 1 O atom \hookrightarrow Produce H_2O Dioxygenase \rightsquigarrow 2 O atoms \hookrightarrow X H_2O	Peroxidase Oxidation of substrate using H_2O_2 as reactants ☆ Glutathione peroxidase \hookrightarrow Protect the body from ROS $2 \text{ Glut} \xrightleftharpoons[\text{Reductase NADP}]{\text{Peroxidase}} \text{Glut}$

2) Transferase

Kinase Transfer a phosphate from (ATP, GTP, ...) to a substrate such as: phosphofruktokinase $\text{Fructose 6-phosphate} \longrightarrow \text{Fructose 1,6 bisphosphate}$	Transaminase Transfer amino group Convert amino acids \rightleftharpoons keto acids Aspartate \rightleftharpoons Oxaloacetate Alanine \rightleftharpoons Pyruvate Glutamate \rightleftharpoons α -keto glutarate	Synthase Transfer monomers to build polymers Such as Glycogen synthase
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3) **Hydrolase** \rightsquigarrow Cleavage reactions using water
 \hookrightarrow such as protease, lipase, Glycosidase

4) Lyase \rightsquigarrow Cleavage without using water

Dehydrases

Remove H_2O (product)
form double bond
Such as enolase

Decarboxylase

Remove Carboxyl
produce CO_2

Synthase

Citrate synthase

Aldolase

break fructose 1,6-bisphosphate
into Glyceraldehyde 3-phosphate
and Dihydroxyacetone phosphate

5) Isomerase

Isomerase \rightsquigarrow Rearrange bonds

Mutase \rightsquigarrow move phosphate

6) Ligase (synthetase)

Join molecules using ATP, GTP, ...

such as Carboxylase

Abzymes

\rightsquigarrow antibodies used as enzymes, produced against transition state analogs

such as abzymes similar to Cocaine esterase

Ribozymes

\rightsquigarrow Non protein enzymes (RNA + protein)

catalysis

\rightsquigarrow RNA splicing

and protein synthesis

\rightsquigarrow stabilize and enhance the activity of RNA