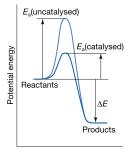
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Reaction coordinate

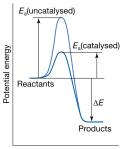
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Reaction coordinate

Homgoeneous: catalyst in same phase as reaction mixture

e.g.,
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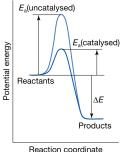


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metal provides surface upon which reactants bind binding facilitates encounters between reactants and increases rxn rate



 $Br^- catalysed \ decomposition \ of \ H_2O_2: \ 2H_2O_2(aq) \longrightarrow 2H_2O(l) + O_2(g)$

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rate:
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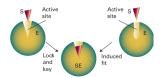
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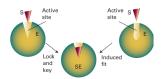
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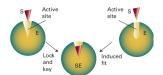
lock-n-key: active site and substrate have complementary 3D structures





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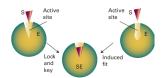
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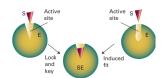


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Many enzymes consist primarily of proteins some featuring organic or inorganic co-factors in active sites



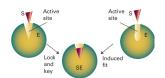
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Certain RNA molecules can also be biological catalysts, forming ribozymes example of ribozyme \Rightarrow ribosome, large assembly of proteins and catalytically active RNA molecules responsible for synthesis of proteins in cell

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- 3. For a given $[E]_0$ and high values of $[S]_0$, rate of product formation : independent of $[S]_0$ reaches maximum velocity, ν_{max}

Michaelis-Menten mechanism: $E+S \underset{k_a}{\overset{k_a}{\rightleftharpoons}} ES \xrightarrow{k_b} P+E$;

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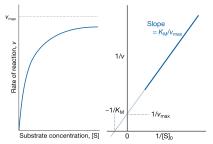
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$$[E] = [E]_0 - [ES];$$
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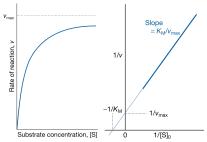


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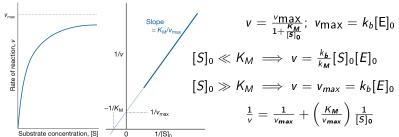
$$v = rac{v_{ ext{max}}}{1 + rac{K_{ ext{M}}}{|S|_0}}; \ v_{ ext{max}} = k_b[\mathsf{E}]_0$$

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$$v_{max} = \frac{v_{max}}{v_{max}}; \quad v_{max} = k_b[E]_0$$

$$[S]_0 \ll K_M \Longrightarrow v = v_{max} = k_b[E]_0$$

$$[S]_0 \gg K_M \Longrightarrow v = v_{max} = k_b[E]_0$$

 $-1/K_{N}$

Substrate concentration, [S]

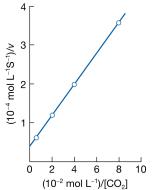
1/[S]₀

 $\frac{1}{V} = \frac{1}{V_{max}} + \left(\frac{K_M}{V_{max}}\right) \frac{1}{[S]_0}$

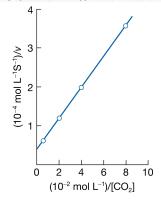
Lineweaver-Burk plot

Lineweaver-Burk plot

Lineweaver-Burk plot catalysis of hydration of CO_2 in red blood cells by carbonic anhydrase $[CO_2]/(mmol dm^{-3})$ 1.25 2.5 5 20



Lineweaver-Burk plot catalysis of hydration of CO_2 in red blood cells by carbonic anhydrase



$$v_{max} = \frac{1}{4} = 0.25 \, \text{mmol.dm}^{-3} \, \text{s}^{-1}$$

Lineweaver-Burk plot catalysis of hydration of CO₂ in red blood cells by carbonic anhydrase

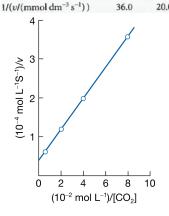
[CO₂]/(mmol dm⁻³) 1.25 2.5 5 20 rate/(mmol dm⁻³ s⁻¹) 2.78×10^{-2} 5.00×10^{-2} 8.33×10^{-2} 1.67×10^{-1} 1/([CO₂]/(mmol dm⁻³)) 0.800 0.200 0.4000.0500

20.0

12.0

6.0

36.0

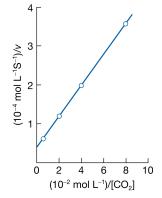


$$v_{max} = \frac{1}{4} = 0.25 \,\mathrm{mmol.dm^{-3}s^{-1}}$$

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Lineweaver-Burk plot

catalysis of hydration of CO_2 in red blood cells by carbonic anhydrase



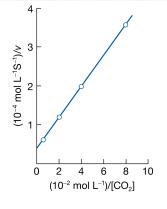
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$$k_{cat} = \frac{v_{max}}{[E]_0} = \frac{2.5 \times 10^{-4}}{2.3 \times 10^{-9}} = 1.1 \times 10^5 s^{-1}$$

Lineweaver-Burk plot

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$$\epsilon = \frac{k_{\text{cat}}}{K_{\text{M}}} = \frac{1.1 \times 10^{5}}{10^{-2}} = 1.1 \times 10^{7} \text{dm}^{3} \text{mol}^{-1} \text{s}^{-1}$$

turnover frequency or catalytic constant, $k_{cat} = \frac{\# \text{catalytic cycles performed by active site}}{\text{duration of interval}}$ has units of 1st order rate constant

turnover frequency or catalytic constant, $k_{cat} = \frac{\# \text{catalytic cycles performed by active site}}{\text{duration of interval}}$ has units of 1st order rate constant numerically equivalent to k_b in Michaelis-Menten mechanism $k_{cat} = k_b = \frac{v_{max}}{|E|_0}$

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Higher $\varepsilon \implies$ more efficient

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 \implies rate constants $\approx 10^8 - 10^9 \text{ dm}^3 \text{mol}^{-1} \text{s}^{-1} \text{(catalytic perfection)}$

$$\begin{array}{lll} \mathsf{E} + \mathsf{S} \rightleftharpoons \mathit{ES} & k_a, \ k_a' \\ \mathsf{ES} \rightarrow \mathsf{E} + \mathsf{P} & k_b \\ \mathsf{E} ! \rightleftharpoons \mathsf{E} + \mathsf{I} & \mathcal{K}_I = \frac{[E][I]}{[EI]} \\ \mathsf{ES} ! \rightleftharpoons \mathsf{ES} + \mathsf{I} & \mathcal{K}_I' = \frac{[ES][I]}{[ESI]} \end{array}$$

$$\begin{array}{lll} \mathsf{E} + \mathsf{S} \rightleftharpoons \mathsf{ES} & k_a, \ k_a' \\ \mathsf{ES} \to \mathsf{E} + \mathsf{P} & k_b \\ \mathsf{E} | \rightleftharpoons \mathsf{E} + \mathsf{I} & \mathcal{K}_I = \frac{[E][I]}{[EI]} \\ \mathsf{ESI} \rightleftharpoons \mathsf{ES} + \mathsf{I} & \mathcal{K}_I' = \frac{[ES][I]}{[ESI]} \\ [\mathsf{E}]_0 = [\mathsf{E}] + [\mathsf{EI}] + [\mathsf{ES}] + [\mathsf{ES}] \end{array}$$

define:
$$lpha=1+rac{[l]}{K_{l}}$$
 $lpha'=1+rac{[l]}{K_{l}'}$

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define:
$$\alpha = 1 + \frac{|\mathbb{I}|}{K_I}$$

$$\alpha' = 1 + \frac{|\mathbb{I}|}{K_I'}$$

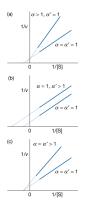
$$[\mathbb{E}]_0 = [\mathbb{E}]\alpha + [\mathbb{ES}]\alpha'$$
using $K_M = \frac{[\mathbb{E}][S]}{[\mathbb{ES}]}$

$$\begin{array}{lll} \mathsf{E} + \mathsf{S} & \mathsf{ES} & k_a, \ k_a' \\ \mathsf{ES} \rightarrow \mathsf{E} + \mathsf{P} & k_b \\ \mathsf{E} | \rightleftharpoons \mathsf{E} + \mathsf{I} & \mathsf{K}_I = \frac{[E][I]}{[EI]} \\ \mathsf{ESI} \rightleftharpoons \mathsf{ES} + \mathsf{I} & \mathsf{K}_I' = \frac{[ES][I]}{[ESI]} \\ [\mathsf{E}]_0 = [\mathsf{E}] + [\mathsf{EI}] + [\mathsf{ES}] + [\mathsf{ES}] \end{array}$$

$$\begin{aligned} \text{define: } \alpha &= 1 + \frac{|I|}{K_I} \\ \alpha' &= 1 + \frac{|I|}{K_I'} \\ \text{[E]}_0 &= [E] \alpha + [ES] \alpha' \\ \text{Using } K_M &= \frac{[E][S]}{[ES]} \\ \text{ES} \rightarrow \text{E+P} & k_b \\ \text{EI} \rightleftharpoons \text{E+I} & K_I &= \frac{[E][I]}{[EI]} \\ \text{ESI} \rightleftharpoons \text{ES+I} & K_I' &= \frac{[ES][I]}{[ESI]} \\ \text{[E]}_0 &= [E] + [EI] + [ES] + [ESI] \end{aligned} \qquad \begin{aligned} \text{define: } \alpha &= 1 + \frac{|I|}{K_I} \\ \text{using } K_M &= \frac{[E][S]}{[ES]} \\ \text{using } K_M &= \frac{[E][S]}{[ES]} \\ \text{[E]}_0 &= \frac{K_M[ES]}{[S]_0} \alpha + [ES] \alpha' &= [ES] \left(\frac{\alpha K_M}{[S]_0} + \alpha'\right) \\ v &= k_b [ES] &= \frac{k_b [E]_0}{\alpha K_M} + \alpha' &= \frac{v_{max}}{\alpha' + \frac{\alpha K_M}{[S]_0}} \\ \text{or, } \frac{1}{v} &= \frac{\alpha'}{v_{max}} + \alpha \left(\frac{K_M}{v_{max}}\right) \frac{1}{[S]_0} \end{aligned}$$

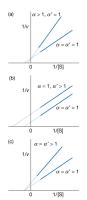
Lineweaver-Burk plots characteristic of three modes of enzyme inhibition:

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Lineweaver-Burk plots characteristic of three modes of enzyme inhibition: competitive inhibition: inhibitor binds only to active site of enzyme and inhibits attachment of substrate; $\alpha>1$ and $\alpha'=1$

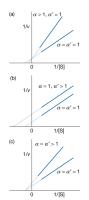
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 $\label{lineweaver-Burk} \mbox{Lineweaver-Burk plots characteristic of three modes of enzyme inhibition:}$

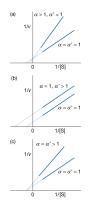
competitive inhibition: inhibitor binds only to active site of enzyme and inhibits attachment of substrate: $\alpha > 1$ and $\alpha' = 1$

uncompetitive inhibition: inhibitor binds to enzyme site removed from active site, but only if substrate is already present. ESI reduces the concentration of ES (active complex). In this case $\alpha = 1$ (: El does not form) and $\alpha' > 1$.

non-competitive (mixed) inhibition: inhibitor binds to site other than active site, and its presence reduces ability of substrate to bind to active site.

Inhibition occurs at both E and ES sites : $\alpha > 1$ and $\alpha' > 1$.

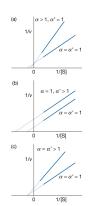
$$\frac{1}{v} = \frac{\alpha'}{v_{max}} + \alpha \left(\frac{K_{M}}{v_{max}}\right) \frac{1}{[S]_{0}}$$



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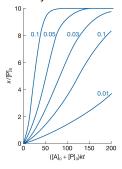


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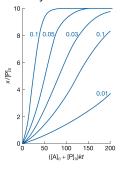
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Slope and y-intercept of Lineweaver-Burk plot increase upon addition of inhibitor. Fig. c : special case : $K_I = K_I'$; and $\alpha = \alpha'$, which results in intersection of lines on x-axis



$$\begin{array}{ccc}
A \longrightarrow P & v = k[A][P] \\
[A]_0 - x & x
\end{array}$$

labelled by b

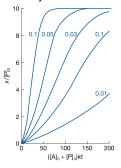


labelled by
$$b$$

$$A \longrightarrow P \qquad v = k[A][P]$$

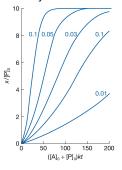
$$[A]_0 - x \quad x$$

$$\frac{dx}{dt} = k([A]_0 - x)([P]_0 + x)$$



labelled by b

$$\begin{split} \mathbf{A} &\longrightarrow \mathbf{P} \qquad \mathbf{v} = \mathbf{k}[\mathbf{A}][\mathbf{P}] \\ [\mathbf{A}]_0 - \mathbf{x} \quad \mathbf{x} \\ \frac{d\mathbf{x}}{dt} &= \mathbf{k} \left([\mathbf{A}]_0 - \mathbf{x} \right) \left([\mathbf{P}]_0 + \mathbf{x} \right) \\ \text{integrating:} \\ \frac{1}{[\mathbf{A}]_0 + [\mathbf{P}]_0} \ln \frac{\left([\mathbf{P}]_0 + \mathbf{x} \right) [\mathbf{A}]_0}{[\mathbf{P}]_0 \left([\mathbf{A}]_0 - \mathbf{x} \right)} = \mathbf{k} \mathbf{t} \end{split}$$

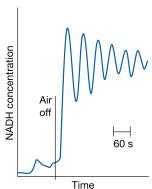


labelled by b

$$\begin{array}{ll} \mathsf{A} \longrightarrow \mathsf{P} & v = k[\mathsf{A}][\mathsf{P}] \\ [\mathsf{A}]_0 - x & x \\ \frac{dx}{dt} = k\left([\mathsf{A}]_0 - x\right)\left([\mathsf{P}]_0 + x\right) \\ \text{integrating:} \\ \frac{1}{[\mathsf{A}]_0 + [\mathsf{P}]_0} \ln \frac{\left([\mathsf{P}]_0 + x\right)[\mathsf{A}]_0}{[\mathsf{P}]_0\left([\mathsf{A}]_0 - x\right)} = kt \\ \\ \mathsf{or,} & \frac{x}{[\mathsf{P}]_0} = \frac{e^{at} - 1}{1 + be^{at}}, \text{ where } a = \left([\mathsf{A}]_0 + [\mathsf{P}]_0\right)k \\ \\ \mathsf{and} & b = \frac{[\mathsf{P}]_0}{[\mathsf{A}]_0} \end{array}$$

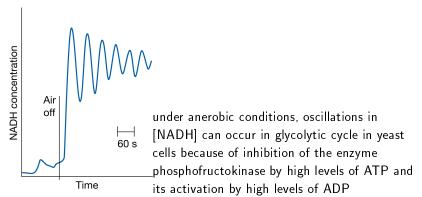
oscillating reactions: Feedback mechanism in which a product either increases o decreases the reaction rate

- can be sustained indefinitely only if the reaction is carried at far from equilibrium conditions (continuous supply and dissipation of species)



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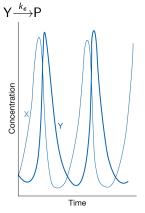
$$A+X \xrightarrow{k_a} 2X; \quad \frac{d[X]}{dt} = k_a[A][X] - k_b[X][Y]$$

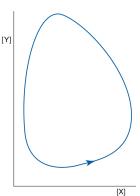
$$A + X \xrightarrow{k_a} 2X; \quad \frac{d[X]}{dt} = k_a[A][X] - k_b[X][Y]$$

$$X+Y \xrightarrow{k_b} 2Y; \quad \frac{d[Y]}{dt} = k_b[X][Y] - k_c[Y]$$

$$\begin{array}{ll} \mathsf{A} + \mathsf{X} \xrightarrow{k_a} 2\mathsf{X}; & \frac{d[X]}{dt} = k_a[A][X] - k_b[X][Y] \\ \mathsf{X} + \mathsf{Y} \xrightarrow{k_b} 2\mathsf{Y}; & \frac{d[Y]}{dt} = k_b[X][Y] - k_c[Y] \end{array}$$

$$X+Y \xrightarrow{\kappa_b} 2Y; \quad \frac{d[Y]}{dt} = k_b[X][Y] - k_c[Y]$$





Belousov-Zhabotinski reaction (KBrO $_3$, malonic acid, cerium (IV) salt in acidic solution):

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 $\mathsf{BrO}_3^-\!+\!\mathsf{HBrO}_2\!+\!\mathsf{H}_3\mathsf{O}^+\longrightarrow\!\!2\mathsf{BrO}_2\!\cdot\!+\!2\mathsf{H}_2\mathsf{O}$

Belousov-Zhabotinski reaction (KBrO₃, malonic acid, cerium (IV) salt in acidic solution):

 $\begin{array}{l} BrO_3^- + HBrO_2 + H_3O^+ \longrightarrow 2BrO_2 \cdot + 2H_2O \\ 2BrO_2 \cdot + 2Ce(III) + 2H_3O^+ \longrightarrow 2HBrO_2 + 2Ce(IV) + 2H_2O \end{array}$

Belousov-Zhabotinski reaction (KBrO₃, malonic acid, cerium (IV) salt in acidic solution):

$$\mathsf{BrO}_3^- + \mathsf{HBrO}_2 + \mathsf{H}_3\mathsf{O}^+ \longrightarrow 2\mathsf{BrO}_2 \cdot + 2\mathsf{H}_2\mathsf{O}$$

 $2BrO_2 + 2Ce(III) + 2H_3O^+ \longrightarrow 2HBrO_2 + 2Ce(IV) + 2H_2O$

product HBrO₂ is a reactant in first step and provides a feedback mechanism that enhances rate of formation of HBrO₂

 $\label{eq:https://en.wikipedia.org/wiki/Brusselator} \begin{array}{l} A \to X \\ 2X + Y \to 3X \\ B + X \to Y + D \\ X \to E \end{array}$

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https://en.wikipedia.org/wiki/Brusselator A \rightarrow X  
2X + Y \rightarrow 3X  
B + X \rightarrow Y + D  
X \rightarrow E  
https://en.wikipedia.org/wiki/Oregonator A + Y \longrightarrow X + P  
X + Y \longrightarrow 2 P  
A + X \longrightarrow 2 X + 2 Z  
A + X \longrightarrow A + B   
A + X \longrightarrow
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