

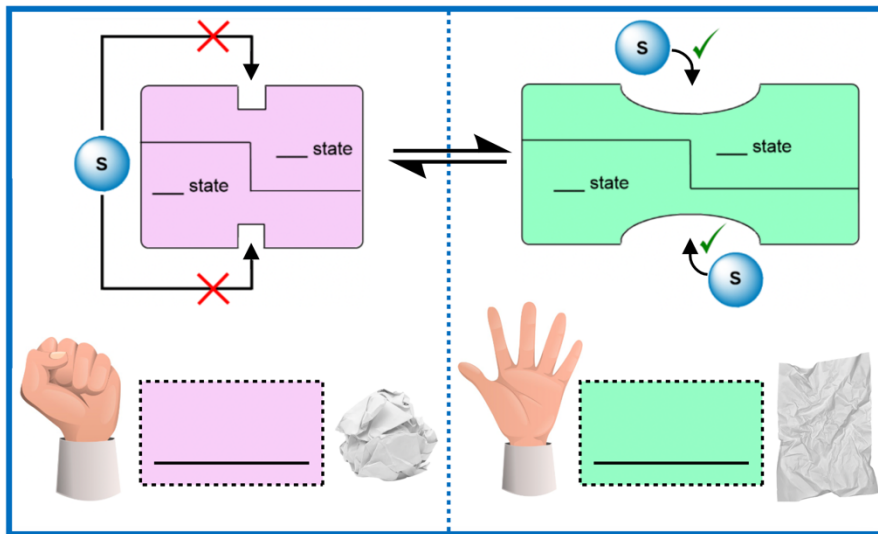
CONCEPT: ALLOSTERIC ENZYME CONFORMATIONS

- Protein _____: alternative 3-dimensional *states* that a protein can achieve.
 - Recall: proteins are not completely rigid structures; protein structures can be induced to changes.
 - Different protein *conformations* can have different abilities and/or _____.

T State & R State

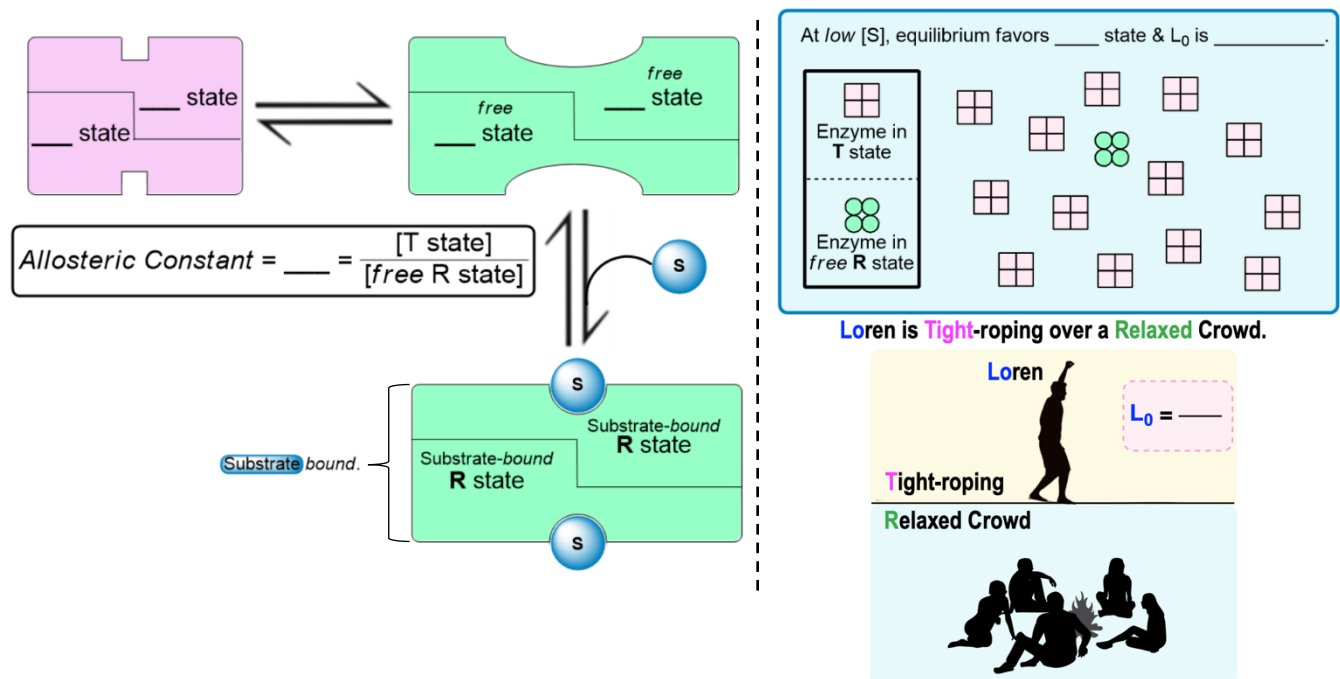
- Allosteric enzymes can exist in one of two states: 1) _____ State (Tense State) 2) _____ State (Relaxed State)
 - T state: catalytically _____ & has a _____ affinity for substrates (binds substrates *inefficiently*).
 - R state: catalytically _____ & has a _____ affinity for substrates (binds substrates *efficiently*).

EXAMPLE:



Allosteric Constant (L_0)

- _____ Constant (L_0): _____ of T States over *free* R States (T/R) when *no substrate* is present.
 - T State is more _____ than *free* R State, so at *low* [S], equilibrium favors _____ State.



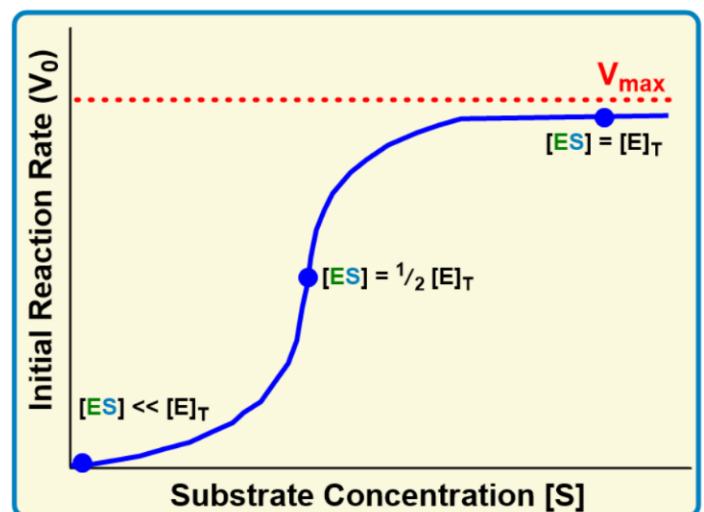
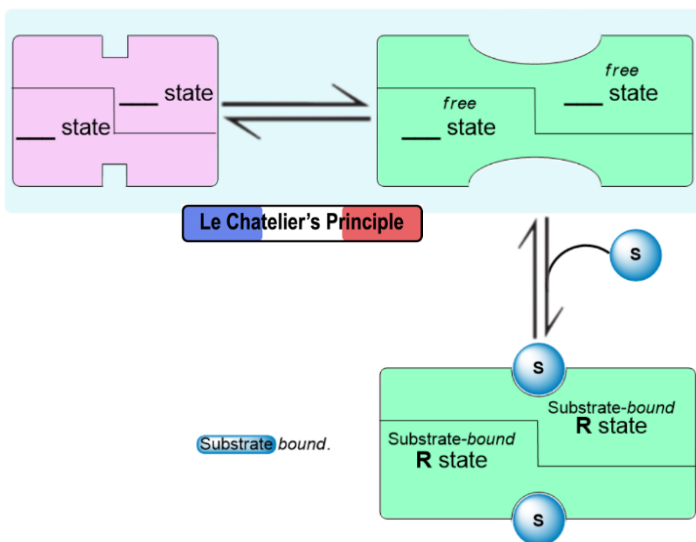
CONCEPT: ALLOSTERIC ENZYME CONFORMATIONS

PRACTICE: Which of the following is true about allosteric enzyme conformational states?

- a) The T state is more stable than the R state of the enzyme when no substrate is present.
- b) Rearrangement of the protein's secondary structure dictates T vs. R states.
- c) The R state of the enzyme has a higher affinity for substrate molecules than the T state.
- d) When a substrate is released from the R state, the enzyme remains in that state indefinitely.
- e) All of the above are correct.
- f) Only A and D are correct.
- g) Only A and C are correct.

T/R Conformations Allow for Cooperative Kinetics

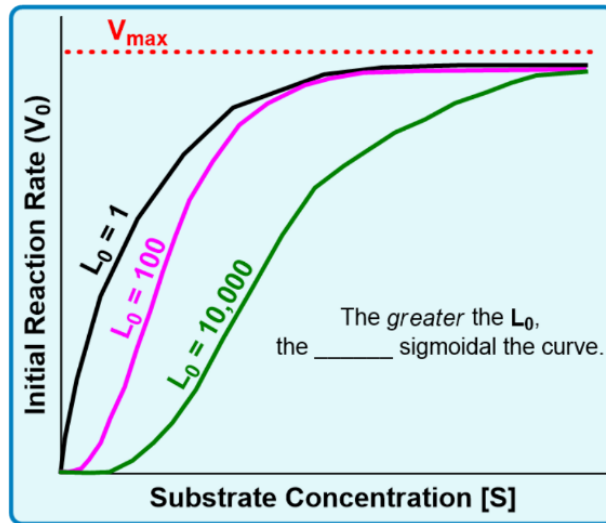
- *Sigmoidal* kinetics ("S"-shaped curve) displayed by *allosteric* enzymes suggests that **S** binding is _____.
 - *Positive Cooperativity*: binding of one **S** molecule makes it _____ for other **S** molecules to bind enzymes.
 - *Question*: How does cooperative **S**-binding work?
- Recall: at *low* [**S**], equilibrium favors ____ State; HOWEVER, increasing [**S**] *disrupts* this equilibrium.
- S binding to *free* R state produces **S**-bound-R-state, but consequently _____ [*free* R state].
 - By Le Chatelier's Principle, lowering [*free* R state] causes reaction to shift towards the *free* ____ state.



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L_0 Dictates Curve in Kinetics Plots

- The allosteric constant (____), dictates the extent of an *allosteric enzyme*'s sigmoidal curve.
 - The *greater* the L_0 , the _____ *sigmoidal* the curve will be in a kinetics plot (V_0 vs $[S]$).
 - The *smaller* the L_0 , the _____ sigmoidal and the more the curve resembles *Michaelis-Menten* kinetics.



- _____ models explain the *sigmoidal* kinetics of *allosteric* enzymes:
 - 1) _____ (or _____) Model.
 - 2) _____ (or _____) Model.
 - In *both* models, allosteric enzyme reaction activity can be affected by allosteric _____.

PRACTICE: An allosteric enzyme that follows the concerted model mechanism has a $L_0 = 10,000$ in the absence of substrate. A mutation in this enzyme caused the L_0 to now be $1/10,000$ (reciprocal to its original value). What affect does this mutation have on the reaction rate of the enzymatic reaction?

- The enzyme will retain the T state and the reaction will not occur.
- Reaction rate remains independent of the substrate concentration.
- The association constant (K_a) for formation of the enzyme-substrate complex will not change with the mutation.
- Kinetics will appear to be similar to Michaelis-Menten kinetics, since the enzyme is nearly always in its R state.