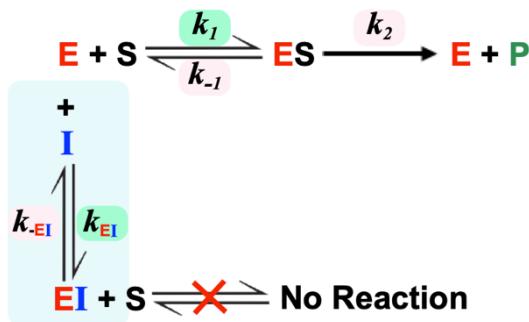


## CONCEPT: DEGREE OF INHIBITION

- **Question:** How can we measure inhibitor effects on the initial rate/velocity ( $V_0$ ) of an enzyme-catalyzed-reaction?
- **\_\_\_\_\_ of inhibition:** a factor that measures/quantifies how much an enzyme is actually being *inhibited* by the inhibitor.
- Recall: inhibitors can *either* bind to the \_\_\_\_\_ enzyme to form EI-complex or to the \_\_\_\_\_-complex to form ESI-complex.
  - Biochemists can *separately* measure the *degree of inhibition* on the \_\_\_\_\_ enzyme & the \_\_\_\_\_-complex.

### $\alpha$ Measures Effects of Inhibition on Free Enzyme (E)

- Degree of Inhibition on \_\_\_\_\_ enzyme (\_\_\_\_) =  $1 + \frac{[I]}{K_I}$ 
  - $\alpha$  is a *unitless* factor that is \_\_\_\_\_ greater than or equal to 1 ( $\alpha \geq 1$ ).
  - No inhibitor is present when  $\alpha = \underline{\hspace{1cm}}$ , AND as  $\alpha$  \_\_\_\_\_, [inhibitor] *increases*.



$$\text{Degree of Inhibition on \_\_\_\_\_\_ Enzyme} = \alpha = \left(1 + \frac{[I]}{K_I}\right)$$

$\alpha < 1$	
$\alpha = 1$	Inhibitor is _____ (____ inhibitor)
$\alpha > 1$	Inhibitor is _____.

**EXAMPLE:** According to the data in the table below, which enzyme is affected most by the inhibitor?

- a) Enzyme A.      b) Enzyme B.      c) Enzyme C.

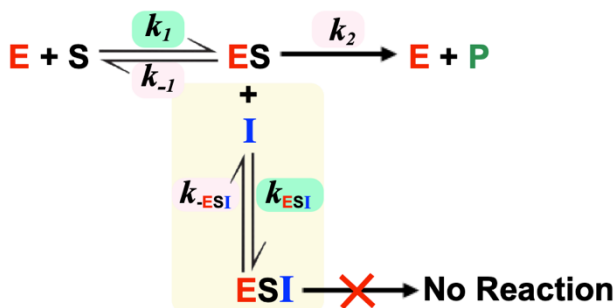
Enzyme A	$\alpha = 1.37$
Enzyme B	$\alpha = 1.98$
Enzyme C	$\alpha = 1.02$

**PRACTICE:** Calculate the degree of inhibition of an inhibitor on the free enzyme if the  $[I] = 3 \mu\text{M}$  and the  $K_I = 6 \mu\text{M}$ .

- a)  $\alpha = 1.5$       b)  $\alpha = 2$       c)  $\alpha = 4$       d)  $\alpha = 3.5$       e)  $\alpha = 0.5$

### $\alpha'$ Measures Effects of Inhibition on ES-Complex

- Degree of Inhibition on \_\_\_\_\_-Complex (\_\_\_\_) quantifies effect of an inhibitor on the ES-complex ( $\alpha'$ ).



$$\text{Degree of Inhibition on \_\_\_\_\_\_-complex} = \alpha' = \left(1 + \frac{[I]}{K'_I}\right)$$

- $\alpha$  and  $\alpha'$  are used as factors to modify the *Michaelis-Menten* & *Lineweaver-Burk* equations in the presence of inhibitors.

**CONCEPT: DEGREE OF INHIBITION**

**PRACTICE:** Calculate the degree of inhibition of an inhibitor on the ES-complex if the  $[I] = 8 \mu\text{M}$  and the  $K'_I = 0.03 \text{ mM}$ .

- |                    |                    |                    |
|--------------------|--------------------|--------------------|
| a) $\alpha = 0.87$ | c) $\alpha = 1.27$ | e) $\alpha = 0.27$ |
| b) $\alpha = 1.08$ | d) $\alpha = 2.39$ |                    |