

(19) **Today**

6.3 Enzyme Kinetics

6.4 Enzyme Inhibition

6.5 An Enzymatic Mechanism

**Next Class (20)**

6.5 An Enzymatic Mechanism

Chap 7: Carbohydrates

(21) **Second Class from Today**

Chap 7: Carbohydrates

**Third Class from Today (22)**

Chap 7: Carbohydrates

**Monday office hours rescheduled to 1:10 to 2:10 from now on.**

Please hand in reworked test 1.

Test 2 postponed to Wednesday, April 2

$$\text{rate} = k_3[\text{ES}]$$

$$[\text{ES}] = k_1/(k_3 + k_2) [\text{E}][\text{S}]$$

$$K_m = (k_3 + k_2)/k_1$$

$$[\text{E}] = [\text{E}]_T - [\text{ES}]$$

$$[\text{ES}] = [\text{E}]_T [\text{S}]/(K_m + [\text{S}])$$

Which finally gets us to the rate law...

$$\text{rate} = \frac{k_3[\text{E}]_T [\text{S}]}{(K_m + [\text{S}])}$$

# Enzyme Kinetics: Compare the Rate law to the Plot

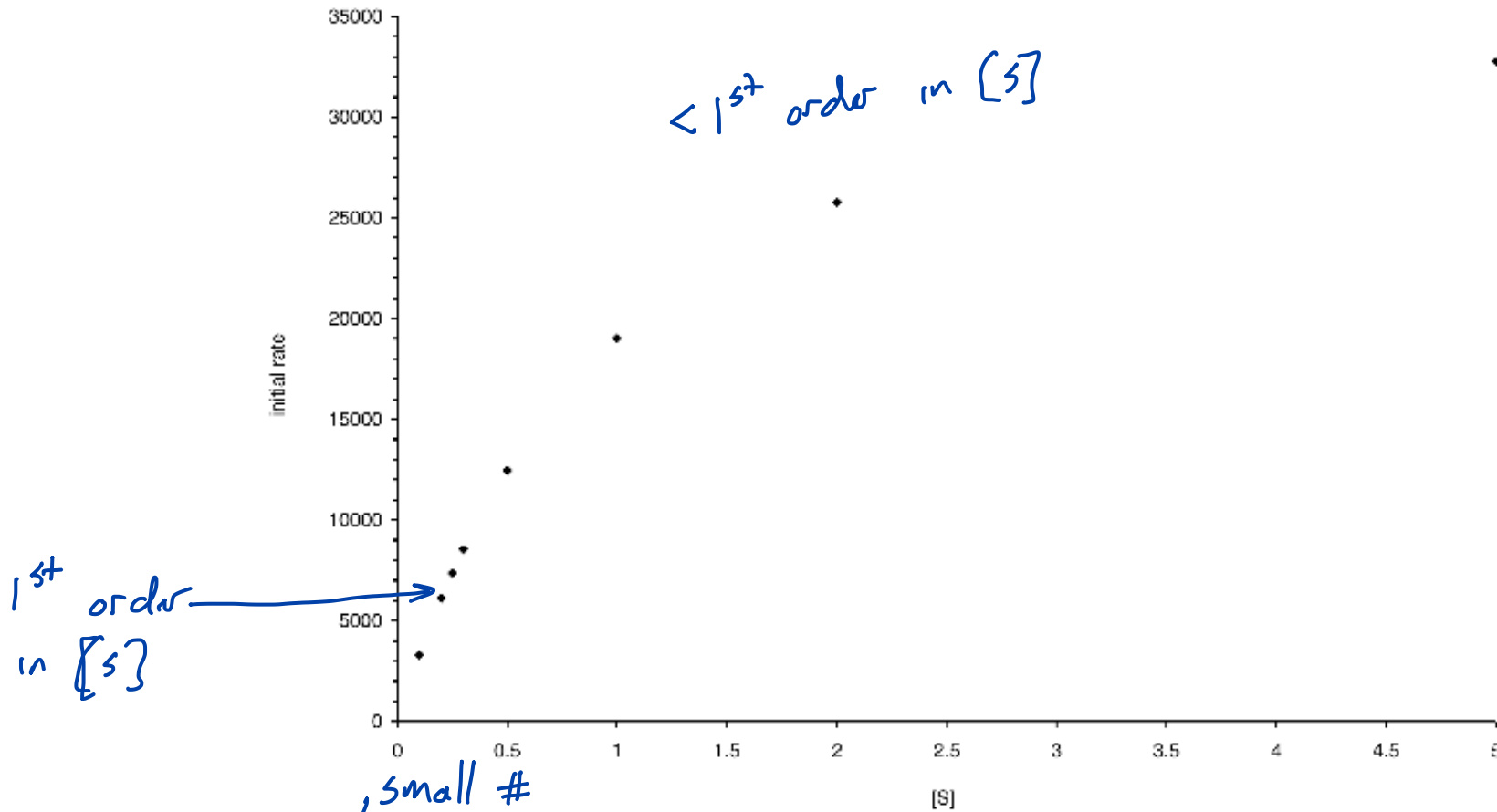
6.3.3

Initial Rate vs Substrate Concentration

0 order in [S]

•  $V_{max}$

$5M = [S]$



$K_m \gg [S]$	$K_m$ similar to $[S]$	$K_m \ll [S]$
$\text{rate} = \frac{k_3[E]_T [S]}{(K_m + [S])}$ <p><math>K_m + [S] \approx K_m</math></p>	$\text{rate} = \frac{k_3[E]_T [S]}{(K_m + [S])}$ <p>less than first order</p>	$\text{rate} = \frac{k_3[E]_T [S]}{[S]}$ <p><math>V_{max}</math></p>

at low  $s$  bottom  $s$  is small so it can be ignored

$$\text{rate} = \frac{k_3[E]_T [S]}{K_m + [S]}$$

$$\frac{1}{\text{rate}} = \frac{K_m + [S]}{k_3[E]_T [S]}$$

$$\frac{1}{\text{rate}} = \frac{K_m}{k_3[E]_T [S]} + \frac{\cancel{[S]}}{k_3[E]_T \cancel{[S]}}$$

$$\frac{1}{\text{rate}} = \left( \frac{K_m}{k_3[E]_T} \right) \frac{1}{[S]} + \frac{1}{k_3[E]_T} \quad \swarrow \quad \frac{1}{v_{\max}}$$

$$y \quad m \quad x \quad + \quad b$$

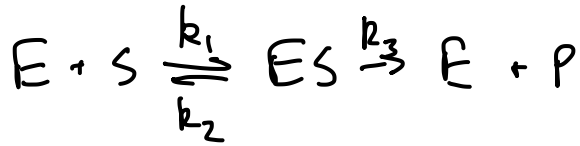
# Enzyme Kinetics: Lineweaver-Burke Plot

6.3.2

$$\text{rate} = \frac{k_3[E]_T [S]}{(K_m + [S])}$$

$$\frac{1}{\text{rate}} = \frac{K_m}{k_3[E]_T} \frac{1}{[S]} + \frac{1}{k_3[E]_T}$$

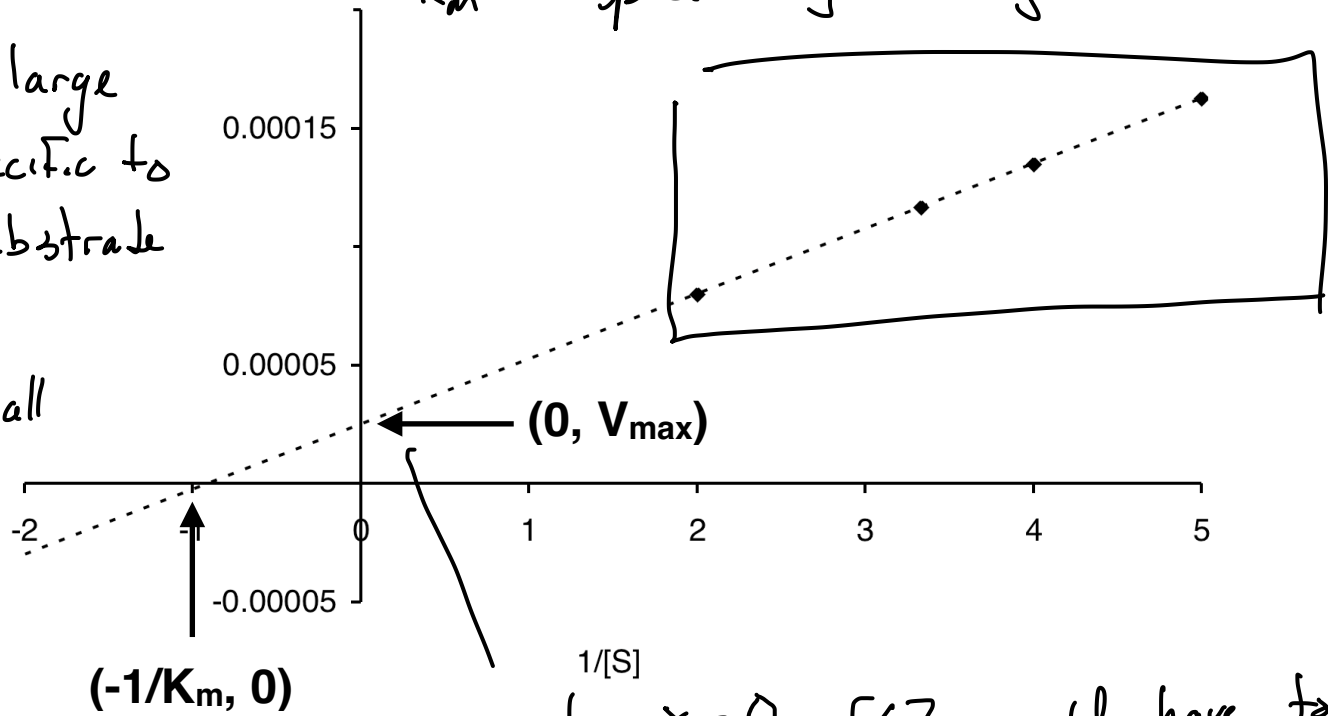
$$V_{\max} = k_3[E]_T$$



$V_{\max}$  intrinsic rate constant ( $k_3$ )  
 $K_m$  specificity for given substrate

$K_m$  small...  $k_1$  large  
 enzyme is specific to  
 particular substrate

$K_m$  large,  $k_1$  small  
 enzyme is less  
 specific



at  $x=0$   $[S]$  would have to be ...

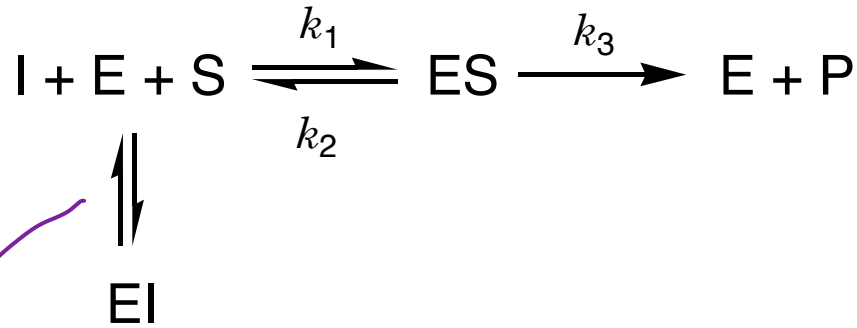
$$[S] = \infty$$

$$\lim_{[S] \rightarrow \infty} \frac{1}{[S]} = 0$$

If  $\frac{k_3 + k_2}{k_1}$  is large  $k_2$  is large  
 $k_1$  and  $k_3$  is small and  
 ES dissociates easily

If  $K_m$  is small  $k_1$  is large  
 large  $k_1$  means large binding constant

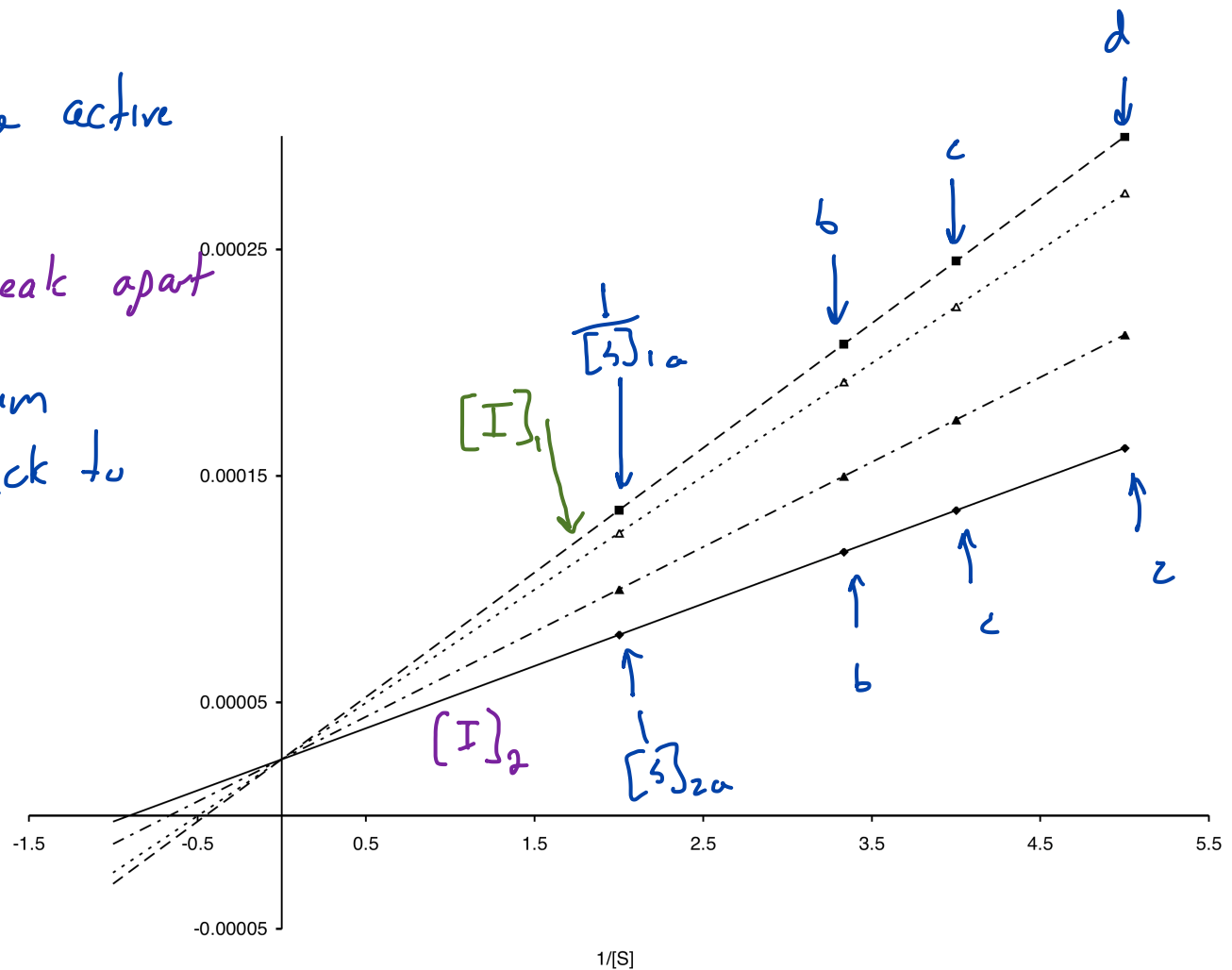
# Competitive Inhibition



$E + I$  compete for the active site

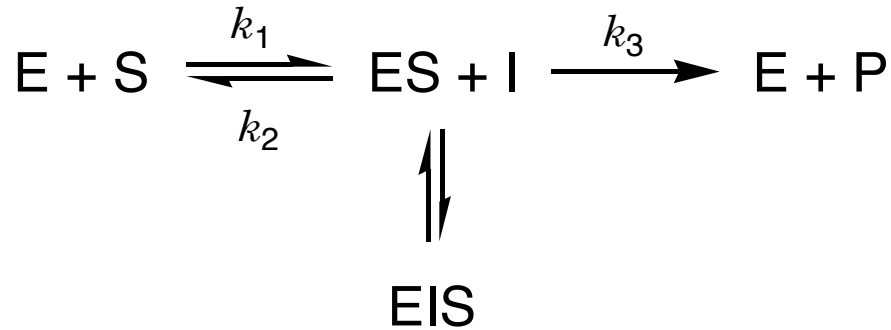
Equilibrium so  $EI$  can break apart

Increase  $[S]$  drives equilibrium shifts  $EI$  equilibrium back to  $E + I + S$  so  $V_{max}$  can still be reached



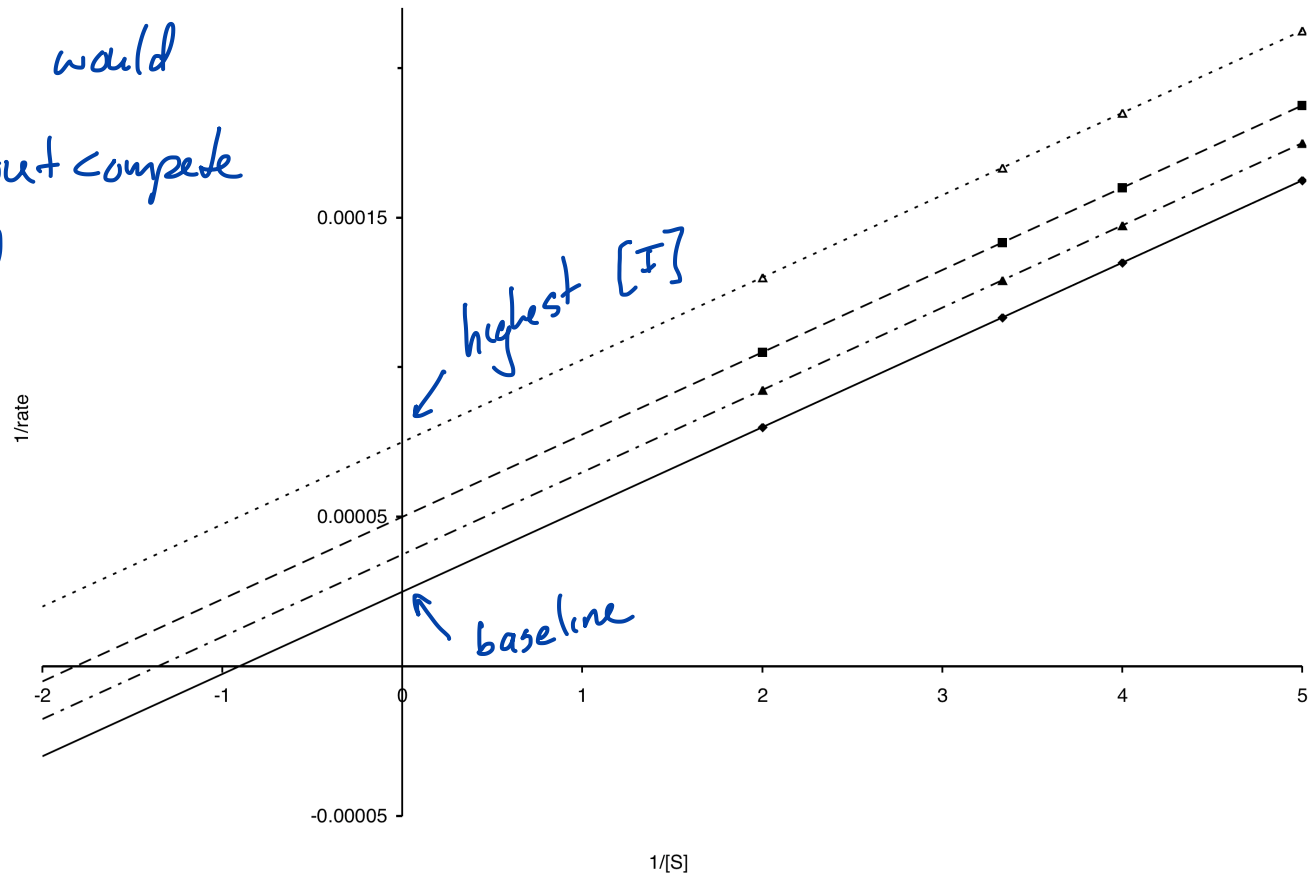
# Uncompetitive Inhibition

6.4.3



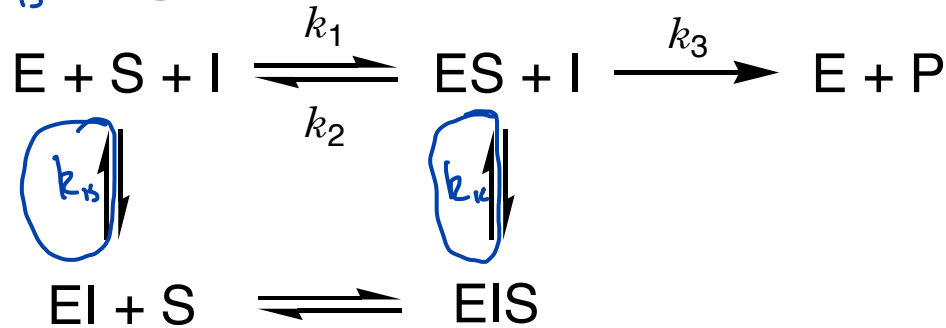
Increase  $[S]$  increases  $[ES]$   
 but if  $[ES]$  increases so would  
 $[EIS]$  so we cannot out compete  
 by adding more  $S$  and  
 $V_{max}$  decreases

$K_m$  and  $V_{max}$  are  
 changing by the same



# Noncompetitive or Mixed

$k_{is} = k_{ic}$



$V_{max}$  changes

lines are not parallel

Design a drug to stop/slow  
a metabolic pathway...  
which would be better...  
competitive or uncompetitive

