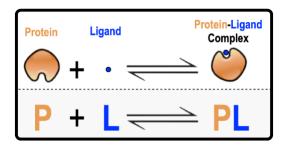
## **CONCEPT:** INTRODUCTION TO PROTEIN-LIGAND INTERACTIONS

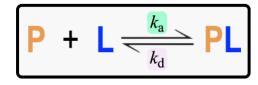
- •\_\_\_\_\_: any small substance that reversibly binds & forms a complex with a larger biomolecule.
  - □ Reversible binding of a protein (\_\_\_\_) to a ligand (\_\_\_\_) is described by a simple expression: P + L ⇒ PL

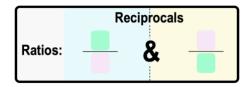


## **Protein-Ligand Rate Constants**

- Recall: every reaction has a \_\_\_\_\_ constant (k) indicating reaction rate efficiency/probability under set conditions.
  - $\Box$  The *higher* the k, the \_\_\_\_\_\_ likely the reaction is \_\_\_\_\_.
- Association Rate Constant (ka): rate constant for free protein & ligand (P + L) \_\_\_\_\_\_ into a complex (PL).
- Dissociation Rate Constant (kd): rate constant for protein-ligand-complex (PL) \_\_\_\_\_\_ back into free P + L.
  - $\Box$  \_\_\_\_\_ of  $k_a$  and  $k_d$  describe the reversible binding of a protein & ligand.

**EXAMPLE:** PL Association & Dissociation Rate Constants:





PRACTICE: Which of the following rate law expressions represents a protein-ligand interaction at equilibrium?

- a)  $[PL]k_a = [P][L]k_d$ .
- b)  $[PL][P]k_a = [L]k_d$ .
- c)  $[L][PL]k_d = [P]k_a$ .
- d) [L][P] $k_a = [PL]k_d$ .

**PRACTICE:** Calculate the dissociation rate constant ( $k_d$ ) at equilibrium if [P] = 20 mM, [L] = 10 mM, [PL] = 5 mM, and the association rate constant ( $k_a$ ) = 100 mM-1s-1.

- a) 400.
- b) 4,000 s-1.
- c) 4,000 mM-1s-1.