**Bio 5312** Problem Set-2 Lohman

1. Consider the equilibrium

## $M + X \Leftrightarrow MX$

If the standard state free energy change is  $\Delta G^{\circ} = -14.2$  kcal/mol for a 1 M standard state at 25°C, for all components what is the standard state free energy change for a 1 mM standard state for all components?

2. Consider the protonation equilibrium

## $M + H \Leftrightarrow MH$

If the standard state free energy change for this reaction is -9.0 kcal/mol for a 1M standard state at 25°C for all components, calculate the free energy change when both M and MH are 1M, but a pH of 7 is maintained?

3. A protein, P, has 10 sites for a ligand, L.

a.) How many different configurations of the protein can occur in which 4 ligands are bound?

b.) If L binds independently and identically to the 10 sites, with site binding constant  $\kappa$ , obtain an expression for the overall macroscopic binding constant,  $\beta_3$  for the equilibrium

$$3L + P \iff PL_3$$

c.) Obtain an expression for the step-wise macroscopic binding constant,  $K_6$ , for the equilibrium:

$$L + PL_5 \iff PL_6$$

d.) For a protein with **n** independent and identical binding sites, what is the value of the ratio of the step-wise macroscopic binding constants,  $K_1/K_n$ ?

4. A homo-tetrameric protein (identical subunits) has a square structure with four binding sites for a ligand X, one site per subunit without cooperativity. X binds to each subunit with a different site binding constant,  $\kappa_i$ . (i.e.,  $\kappa_1$ ,  $\kappa_2$ ,  $\kappa_3$ ,  $\kappa_4$ ).



- a.) Write expressions for the binding polynomial (molecular partition function) and <X> for this tetrahedral system in terms of:
- i. the overall or stoichiometric binding constants,  $\beta_i$  (i.e.,  $\beta_1$ ,  $\beta_2$ ,  $\beta_3$  and  $\beta_4$ ) and the free ligand concentration, *x*.
- ii. the step-wise macroscopic binding constants,  $K_i$  (i.e.,  $K_1$ ,  $K_2$ ,  $K_3$ ,  $K_4$ ) and x.
- iii. the step-wise microscopic binding constant,  $k_i$  (i.e.,  $k_1$ ,  $k_2$ ,  $k_3$ ,  $k_4$ ) and x.
- iv. the site binding constants,  $\kappa_i$  (i.e.,  $\kappa_1$ ,  $\kappa_2$ ,  $\kappa_3$ ,  $\kappa_4$ ) and *x*.
- v. the site binding constants, if all  $\kappa_i = \kappa$ .
- b.) Obtain expressions for the macroscopic constants,  $\beta_i$  and  $K_i$  and the step-wise microscopic constants  $k_i$  in terms of  $\kappa_i$ .