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chapter

5

Protein Function

1. **Relationship between Affinity and Dissociation Constant** Protein A has a binding site for ligand X with a K_d of 10^{-8} M. Protein B has a binding site for ligand X with a K_d of 10^{-6} M. Which protein has a higher affinity for ligand X? Explain your reasoning. Convert the K_d to K_a for both proteins.

Answer: Protein B has a higher affinity for ligand X. The lower K_d indicates that protein B will be half-saturated with bound ligand X at a much lower concentration of X than will protein A. Because $K_a = 1/K_d$, protein A has $K_a = 10^8 \text{ M}^{-1}$; protein B has $K_a = 10^6 \text{ M}^{-1}$.

2. **Negative Cooperativity** Which of the following situations would produce a Hill plot with $n_H = 1.0$? Explain your reasoning in each case.

- (a) The protein has multiple subunits, each with a single ligand-binding site. Binding of ligand to one site decreases the binding affinity of other sites for the ligand.
- (b) The protein is a single polypeptide with two ligand-binding sites, each having a different affinity for the ligand.
- (c) The protein is a single polypeptide with a single ligand-binding site. As purified, the protein preparation is heterogeneous, containing some protein molecules that are partially-denatured and thus have a lower binding affinity for the ligand.

Answer: All three situations would produce $n_H = 1.0$. An n_H (Hill coefficient) of 1.0 generally suggests situation (a)—the classic case of negative cooperativity. However, closer examination of the properties of a protein exhibiting apparent negative cooperativity in ligand binding often reveals situation (b) or (c). When two or more types of ligand-binding sites with different affinities for the ligand are present on the same or different proteins in the same solution, apparent negative cooperativity is observed. In (b), the higher-affinity ligand-binding sites bind the ligand first. As the ligand concentration is increased, binding to the lower-affinity sites produces an $n_H = 1.0$, even though binding to the two ligand-binding sites is completely independent. Even more common is situation (c), in which the protein preparation is heterogeneous. Unsuspected proteolytic digestion by contaminating proteases and partial

Solutions Manual For Principles Of Physical Biochemistry

Hans Kuhn, David H. Waldeck, Horst-Dieter Försterling



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Hailed by advance reviewers as a kinder gentler P Chem text this book meets the needs of an introductory course on physical chemistry and is an ideal choice for courses geared toward pre medical and life sciences students Physical Chemistry for the Chemical and Biological Sciences offers a wealth of applications to biological problems numerous worked examples and around 1000 chapter end problems **Analytical Ultracentrifugation** David Scott, Stephen E Harding, Arther

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Spectroscopic Techniques in Biophysics Giorgio Mario Giacometti, Giovanni Giacometti, 2001 This volume brings together the lectures given during the 1999 session of the School of Pure and Applied Biophysics It concerns a number of spectroscopic tools both experimental and computational frequently encountered in biophysical research The chapters of the book have been compiled from the lecture notes distributed among the participants at the school The authors are specialists in their respective fields **Catalog of Copyright Entries. Third Series** Library of Congress. Copyright Office, 1964

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