## BIRKBECK COLLEGE (University of London)

SCHOOL OF BIOLOGICAL SCIENCES

M.Sc. EXAMINATION FOR INTERNAL STUDENTS ON:

Postgraduate Certificate in Principles of Protein Structure MSc Structural Molecular Biology

CRYS024D7

PRINCIPLES OF PROTEIN STRUCTURE

Thursday 22 August 2013

**Duration of examination: 3 hours** 

10.00 - 13.00

Students will be required to answer <u>10</u> out of 15 questions.

All questions carry 10 marks each.

Each question must start on a <u>new</u> page and the question number written at the top of <u>each</u> sheet.

The exam papers have not been prior-disclosed.

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- 1. Answer all parts;
  - a) Illustrate the CORN law. Which amino acid is exempted from this law? {3 Marks}
  - b) Draw the structures for the side-chains of Tyr, Phe and Trp. Which one is entirely non-polar and what features make the other two polar? {3 Marks}
  - c) What feature of proline causes it to act as an alpha helix breaker? {2 Marks}
  - d) How can serine be modified after initial translation into protein? {2 Marks}
- 2. Answer both parts;
  - a) Draw a Ramachandran plot and indicate the locations occupied by the right-handed alpha helix and beta sheets. {5 Marks}
  - b) How is a peptide bond formed? Indicate where the torsion angles phi, psi and omega are located along the polypeptide backbone? {5 Marks}
- 3. Answer both parts;
  - a) Show schematically the features of an alpha helix. {2 Marks}. How can water distort this structure? {2 Marks}
  - b) What distinguishes a parallel from an anti-parallel beta-sheet? {4 Marks}
  - c) Show schematically how the four-strands in a parallel beta-sheet form a twist. {2 Marks}
- 4. Explain the ridges and grooves model of helix packing. {10 Marks}

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- 5. Answer both parts;
  - a) What is meant by scientific fraud and plagiarism? {5 Marks}
  - b) What points should you consider if you wish to publish a scientific paper. {5 Marks}
- 6. Answer all parts;
  - a) Define the term protein superfold and name three examples. {2 Marks}
  - b) Describe or draw the topology of one type of superfold. {2 Marks}
  - c) Name a specific example of a cytoplasmic protein that is classified as a superfold and give its function. {3 Marks}
  - d) What are the common themes used in the construction of extracellular proteins, and why? {3 Marks}
- 7. Explain the role of gene duplication in the evolution of protein structure. {10 Marks}
- 8. Answer both parts;

You are given a protein sequence and told that it comes from the bacterium *Mycobacterium tuberculosis*. Explain how you could use databases and bioinformatics tools to investigate the following;

- a) The three-dimensional structure or fold of the protein {4 Marks}
- b) Whether the protein is likely to be a good target for the design of drugs against tuberculosis. {6 Marks}
- 9. Discuss how helical and icosahedral symmetries have been used to construct viruses. {10 Marks}

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- 10. Describe the distinctive roles of transcription factors and heat shock proteins in the regulation of the protein life cycle. {10 Marks}
- 11. Answer all parts;
  - a) Define a hydrogen bond {2 Marks}
  - b) Name one amino acid with a side chain that can act as a hydrogen bond donor; one with a side chain that can act as an acceptor; and one with a side chain that can act as either or both. {3 Marks}
  - c) Explain briefly how changes in hydrogen bonding patterns when hydrophobic molecules are dissolved in water can cause linear protein chains to fold into compact structures. {5 Marks}
- 12. Answer all parts;
  - a) Name the six types of reaction used to classify enzymes. {1.5 Marks}
  - b) Describe how a coenzyme can contribute to a reaction mechanism. {2.5 Marks}
  - c) Name and describe the active site of a type of protease. {3 Marks}
  - d) Give an example of a how protease inhibitor has been used to design a drug. {3 Marks}
- 13. Answer both parts;
  - a) Explain what is meant by the term "molecular machine" and describe some of the particular challenges in determining the structures of molecular machines. {3 Marks}
  - b) Using diagrams if you prefer, describe in detail the structure of the enzyme ATP synthase, which has been described as a molecular machine. Give the function of this enzyme and explain its mechanism of action starting from its structure. {7 Marks}

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- 14. Answer all parts;
  - a) Draw a schematic diagram of the structure of a G-protein coupled receptor (GPCR) and indicate;
    - i. The number and nature of the membrane-spanning regions. {2 Marks}
    - ii. The positions of the N and C termini. {2 Marks}
    - iii. The location of the binding sites for the ligand and G protein. {2 Marks}
  - b) Describe the mechanism through which a photon of light will activate the GPCR visual rhodopsin. {4 Marks}
- 15. Answer both parts;
  - a) Describe in detail the structure of a T-cell receptor. How does its structure differ from that of an immunoglobulin? {4 Marks}
  - b) Name the other molecules that are involved in the complex formed by this T-cell receptor that leads to the activation of the T cells. Describe or draw the structure of this complex. {6 Marks}

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