PRIMARY EXAMINATION

BIOCHEMISTRY

Tuesday, 4 December 2001 Time allowed: Two hours

INSTRUCTIONS TO CANDIDATES Question 1 in Section A is compulsory. Answer **three** of the questions in Section B, including one from questions 5 & 6. Use diagrams and formulae wherever possible. Maximum marks: 80.

Please answer each of your four questions in a separate book.

Section A (20 marks) This question is compulsory.

- 1. Write short explanatory answers to **THREE** of the following questions:
 - (a) Describe the mechanism which causes oxyhaemoglobin to have a sigmoidal oxygen dissociation curve allowing tissues to be well supplied with oxygen under a variety of metabolic demands.
 - (b) A medically healthy patient announces that he is in the middle of a 72 hour charity fast. Explain what his metabolic state would be. Would it be safe to proceed with simple dental procedures?
 - (c) Another patient with a science background is concerned about the reactive oxygen species (ROS) which can be produced by a dental x-ray. Reassure her by describing the mechanisms which tissues use to protect themselves from these molecules.
 - (d) Saliva contains a variety of molecules which playa vital role in preserving dental health. Discuss three of these molecules.
 - (e) Which lipoprotein is considered protective against heart disease? Justify your answer by describing the biological function of this class of lipoproteins.

Page 1 of 2.

TURN THE PAGE OVER FOR SECTION B.

Section B (60 marks) Answer THREE of the following FIVE questions. This MUST include at least one from questions 5 and 6

- 2. Describe the structure, properties and biosynthesis of collagen, the main protein found in dentine. Describe briefly one of the other (non-collagenous) dentine proteins.
- 3. Methotrexate (MTX) is a potent inhibitor of the enzyme dihydrofolate reductase and is used in combination chemotherapy for the treatment of acute lymphoblastic leukaemia and breast cancer.

DHF + NADPH + H+ H THF + NADP+

Write notes on 5 of the following 6 points.

- (a) How does MTX enter human cells and why is it retained inside?
- (b) Why does MTX bind more tightly to the target enzyme, dihydrofolate reductase, than does the natural substrate, dihydrofolate?
- (c) When MTX binds to dihydrofolate reductase, is the immediate effect accumulation of DHF or depletion of THF, or both? Provide a reason for your answer-
- (d) Which reactions of purine and pyrimidine biosyntheses are inhibited in cells treated with MTX?
- (e) What are the effects on the pools of deoxynucleotides (dNTPs)?
- (f) Why do cells treated with MTX die?
- 4. Nutrition guidelines are put forward by various national health organisations. These may differ slightly between countries but the main messages are consistent. List 4 examples of dietary guidelines and explain the scientific basis behind each of them.
- 5. It is now possible to chemically synthesis short pieces of DNA of around 20 nucleotides that can be used as "primers" and "probes". Discuss the uses of these reagents.
- 6. What are monoclonal and polyclonal antibodies and what are their uses?

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