University of London

## EXAMINATION FOR INTERNAL STUDENTS

For The Following Qualifications:-
B.Sc. M.Sci.

Mathematics 2503: Introductory Mathematical Biology

COURSE CODE : MATH2503

UNIT VALUE : 0.50

DATE : 22-MAY-06

TIME : $\mathbf{1 0 . 0 0}$

TIME ALLOWED : 2 Hours

All questions may be attempted but only marks obtained on the best four solutions will count.
The use of an electronic calculator is not permitted in this examination.

1. Consider the chemical reaction scheme in which $X$ converts to $Y$ at rate $k_{1}, Y$ converts back to $X$ at rate $k_{-1}$ and also forward to the product $P$ :

$$
X \underset{k_{-1}}{\stackrel{k_{1}}{\rightleftharpoons}} Y \xrightarrow{k_{2}} P
$$

Let $x(t), y(t)$ be the concentrations of $X, Y$ at time $t$.
(a) Write down the differential equations for the evolution of $x$ and $y$ and show that they can be written in the form $\frac{d \mathbf{x}}{d t}=M \mathbf{x}$ where $\mathbf{x}=(x, y)^{T}$ and $M$ is a matrix which you should find.
(b) Now suppose that $k_{1}=2, k_{-1}=1$ and $k_{2}=1$. Find an explicit expression for the evolution of the concentrations of $X, Y$ given that initially $X$ has concentration $x_{0}$ and $Y$ has concentration $y_{0}$.
(c) Sketch the phase plane (i.e. solution curves in ( $x, y$ )-space) to indicate what happens to solutions $(x(t), y(t))$ as $t \rightarrow \infty$ for different initial conditions.
2. An ion channel model has 2 gates, one at each end, that may be either open or closed so that it has 3 states: state 1 , the open-open state; state 2 , the open-closed state; state 3 , the closed-open state. The closed-closed state is to be ignored. The probability per unit time of a channel moving from state $i$ to state $j$ is denoted by $k_{i j}$ for $i, j=1,2,3$. There are no transitions from a state $i$ to state $i$ for $i=1,2,3$. Let $\mathbf{p}(t)=\left(p_{1}(t), p_{2}(t), p_{3}(t)\right)$ be the probability distribution of the channel states at time $t$.
(a) Write down a Markov chain model for the evolution of $\mathbf{p}(t)$.
(b) Suppose that $k_{12}=k_{13}=k_{21}=k_{23}=k_{31}=k_{32}=1$. State what property of the chain ensures that it converges to a unique steady state and find the steady state.
(c) Find the long term probability distribution of the channel states if $k_{13}=k_{21}=$ $k_{23}=k_{31}=1$, but $k_{12}=k_{32}=0$.
3. Suppose $M$ is an invertible real $n \times n$ matrix. Consider the initial value problem

$$
\begin{equation*}
\frac{d \mathbf{x}}{d t}=M \mathbf{x}+\mathbf{a}, t \geqslant 0, \mathbf{x}(0)=\mathbf{x}_{0} \in \mathbb{R}^{n} \tag{1}
\end{equation*}
$$

(a) Find an explicit expression for the solution $\mathbf{x}(t)$ of (1).
(b) Suppose that the eigenvalues of $M$ are real. Briefly discuss how the long term behaviour of the solution $\mathbf{x}(t)$ depends upon the eigenvalues of $M$.

A drug accumulates in the blood at constant rate $\gamma\left(\mathrm{s}^{-1} \mathrm{~mol}^{-1}\right)$. From the blood it transfers to the tissue at rate $\alpha\left(\mathrm{s}^{-1}\right)$ and then from the tissue it transfers to the liver at rate $\beta\left(\mathrm{s}^{-1}\right)$ where it is removed at rate $\delta\left(\mathrm{s}^{-1}\right)$.
(c) Denoting the concentration of the drug in the blood, tissue and liver by $B(t)$, $T(t), L(t)$ respectively, write down a differential equation model for the drug transfer.
(d) Find the limiting concentrations $B(\infty), T(\infty), L(\infty)$. [You are not required to give expressions for $B(t), T(t), L(t)]$.
4. Consider the one-dimensional passive diffusion of a chemical $X$ through a membrane of thickness $h$ spanning from $x=0$ to $x=h$, and with diffusive constant $D$. For $x \leqslant 0$ the concentration of $X$ is maintained at the constant value $C_{0}$ and for $x \geqslant h$ it is maintained at the constant value $C_{1}$.
(a) Write down the partial differential equation for the evolution of the concentration $C(x, t)$ of $X$ in the interval $x \in[0, h]$, together with the model boundary conditions.
(b) Suppose that the concentration of $X$ within the membrane is in steady state. Find the profile of the concentration of $X$ within the membrane.

Now suppose that there are two membranes, one spanning the interval $[0, h)$ with diffusive constant $D_{1}$ and the second spanning the interval $[h, 2 h]$ with diffusive constant $D_{2}$. The concentration of $X$ for $x \leqslant 0$ is maintained at the constant value $C_{0}$ and the concentration of $X$ for $x \geqslant 2 h$ is maintained at the constant value $C_{1}$.
(c) Find the stationary profile of the concentration of $X$ within the two membranes.
5. Consider the system of 3 blood vessels in which vessel 1 bifurcates into vessel 2 and vessel 3 , as shown in figure 1 . For $i=1,2,3$ let $Q_{i}$ denote the flux in vessel $i$, and $P_{i}$ the fixed pressure at the end of vessel $i$. Vessel 1 has length 2 units and vessels 2


Figure 1:
and 3 have unit length. Ohm's law for each vessel is $\Delta P=\alpha L Q$ where $\Delta P$ is the pressure difference across the vessel, $Q$ is the flux, $L$ is the vessel length, and $\alpha>0$ is a constant (same for all vessels).
(a) Find the fluxes $Q_{1}, Q_{2}, Q_{3}$ in terms of the pressures $P_{1}, P_{2}, P_{3}$ and $\alpha$.

Now suppose that Ohm's law for vessel 1 is replaced by the new law $\Delta P=\alpha L|Q| Q$ but remains $\Delta P=\alpha L Q$ for vessels 2 and 3 .
(b) Show that the new flux $Q_{1}$ satisfies

$$
\left|Q_{1}\right| Q_{1}+\frac{Q_{1}}{4}+\frac{1}{4 \alpha}\left(P_{3}+P_{2}-2 P_{1}\right)=0
$$

(c) For fixed pressures $P_{1}>P_{2}$ and $P_{1}>P_{3}$ how many possible fluxes $Q_{1}$ are there? Justify your answer.

