

Mathematical Physiology

Ian Griffiths

Problem Sheets

- Problem Sheet classes:
 - Tuesdays 2–3:30, Weeks 2, 4, 6, 8, Room C5.
Tutor: Sam Palmer. TA: Siting Miao
 - Wednesday 11-12:30, Weeks 2, 4, 6, 8, Room C6.
Tutor: Ian Griffiths. TA: Siting Miao
- Hand-in time for both classes: Friday 5pm of Weeks 1, 3, 5, 7.
- The first problem sheet will be a little shorter since you only have a week to attempt this.

Special Topics

- For those who are attending and need to write a special topic on this course, I have uploaded a list of possible topics.

Lectures

- The typeset lecture notes are [detailed](#). But, [everything you need](#) will be covered in the lectures. (Sometimes I will point to the lecture notes for an additional proof.)
- The course is a little different to other mathematical courses. Here there will be just as much emphasis on coming up with the appropriate mathematical models as there is on solving them.
- To add a little interest and relevance, we will have some guest appearances from research experts in the field (in brain modelling, calcium dynamics,...).
- I will use a mixture of Powerpoint slides (which will be available on the course website) and writing on the whiteboard.

Enzyme kinetics

- Enzymes are catalysts – they help convert other molecules (called substrates) into products but are not used up in the reaction themselves.
- Consider chemicals A and B reacting on collision to form chemical C with a rate k . (The rate depends on the molecule shapes and sizes and the temperature.)

Transmembrane ion transport

- Cells are effectively bags of water.
- The water contains dissolved salts: NaCl and KCl, which dissolve into Na⁺, K⁺ and Cl⁻ ions.
- They exist both inside and outside the cell, creating a potential difference.
- The cell walls are permeable – ions may be transported through the cell membrane, passing through pores called **channels** or **gates**.
- **Osmosis** is the mechanism by which water is transported across the cell membrane.

- **Carrier mediated diffusion** – a molecule hitches a lift by binding to a carrier molecule that is lipid soluble and can move through the membrane.
- **Carrier mediated transport** – a molecule binds to a protein that has an active site that may be exposed to the interior or exterior of the cell (e.g., glucose or amino acid transport).
- **Pumps** – these exchange one ion for another, e.g., Na^+ for K^+ or Na^+ for Ca^{2+} .

Summary of lecture 1

- The law of mass action:



- Michaelis–Menten law:

$$c = \frac{s}{s + K'} \qquad \frac{ds}{dt'} = -\frac{\lambda s}{s + K'}$$

PhD studentship on modelling and data analysis of in-vitro fertilization

- Based at Cardiff Mathematics and supervised by Dr Katerina Kaouri (lead supervisor), Dr Thomas Woolley, Prof Karl Swann (Cardiff Biosciences), Prof Krasimira Tsaneva-Atanaseva (Exeter) and Dr Cameron Hall (Bristol).
- The student should have a strong quantitative background (degree in Mathematics, Physics, Engineering or Computer Science).
- Open to both home and international students.
- The funding is for four years.
- Deadline: **2nd November**. Start date: October 2023
- More info and how to apply: www.findaphd.com/phds/project/calcium-signalling-in-in-vitro-fertilization-developing-a-non-invasive-diagnostic-tool/?p147305
- Contact: kaourik@cardiff.ac.uk

Transmembrane ion transport

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A model for carrier mediated transport

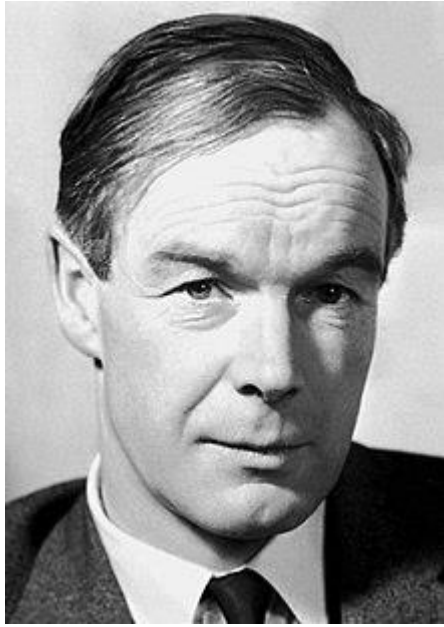
- C_i = a state with a binding site exposed to the interior.
- C_e = a state with a binding site exposed to the exterior.
- C_e can bind with a substrate molecule in the exterior S_e to make a product P_e .
- C_i can bind with a substrate molecule in the interior S_i to make a product P_i (with the same rates as the exterior).
- P_i can turn into P_e . This is the carrier doing its 'rotation'.
- C_i can turn into C_e . This is the carrier site rotating without any substrate on it. We assume this occurs at the same rate as the rotation with substrate on it.

Summary of lecture 2

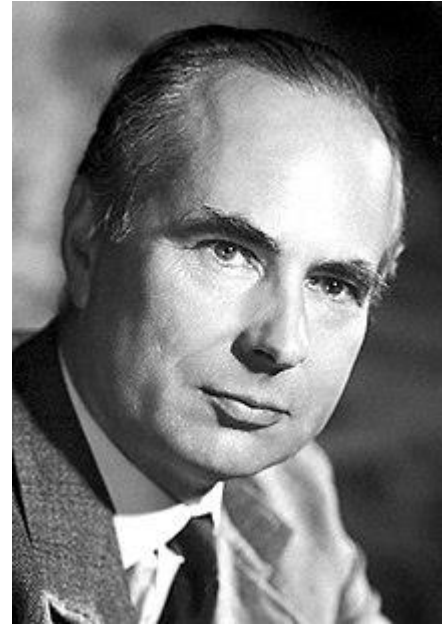
- We can write down an ODE model for carrier-mediated transport. The model tells us the rate at which ions can be transferred from the outside of the membrane to the inside in steady state.
- Active transport involves moving molecules against concentration gradients. This requires energy.
- Ions will move across a membrane wall to balance concentration. However, this might lead to a difference in charge. The system reaches an equilibrium when the diffusive flux balances the ionic flux.
- This occurs at the Nernst potential:

$$V_S = \frac{RT}{zF} \ln \left(\frac{c_e}{c_i} \right)$$

The Hodgkin–Huxley model



1914–1998



1917–2012

The Hodgkin–Huxley model

- The nervous system is a communication systems formed by nerve cells or **neurons**.
- Information is propagated along long cylindrical segments called **axons** by electrochemical signals.
- Communication between cells occurs at junctions between **synapses** to the **dendrites**.
- If a small current is applied for a short time the membrane potential simply returns to its resting potential when the current is removes.
- But for a sufficiently high current, the membrane potential undergoes a large excursion – an **action potential** – before returning to its resting value.
- Signals are transmitted by the propagation of these action potentials.