# MD1A4-45 Integrated Science Organelles and Cells

#### 23/24

Department Warwick Medical School Level Undergraduate Level 1 Module leader Masanori Mishima Credit value 45 Module duration 10 weeks Assessment 70% coursework, 30% exam Study location University of Warwick main campus, Coventry

## Description

## Introductory description

The module aims to equip students with the conceptual, theoretical and computational skills required for the analysis and engineering of prokaryotic and eukaryotic organelles and cells.

Module web page

#### Module aims

Students will learn to solve scientific problems in this area by integrating concepts and approaches from different scientific disciplines, including biology, physics, chemistry, mathematics and computing.

## **Outline syllabus**

This is an indicative module outline only to give an indication of the sort of topics that may be covered. Actual sessions held may differ.

Block 6 Mathematical Skills

Lecture 1 - Basic Statistics

Mean, median, mode, histogram, R, variance, standard deviation, standard error, confidence interval.

Lecture 2 - Trigonometry

Pi, radian, sin, cos, tan, sum & difference identities, sum to product identities

Lecture 3 - Calculus

Integration by substitution, integration by parts

Lecture 4 - ODE

Overview, general solution, particular solution

Lecture 5 - Linear Algebra

Vector, dot product.

Lecture 6 - Statistics

Probability, probability mass finction, probability density function

Lecture 7 & 8 - Coach writeup

Dry Lab 1 - Formula handling

Dry Lab 2 - Calculus

Dry Lab 3 - Complex numbers, waves

Dry Lab 4 - ODE

Dry Lab 5 - Linear algebra

Dry Lab 6 - Statistics

In-class exam

#### **B7** BioElectricity

How does life harness electrical forces?

We will learn how life harnesses electricity to communicate and compute information. We will discover how cells use energy to create electro-chemical gradients and express proteins that behave as selective pores in the plasma membrane to control where, when and how certain ions pass into and out of the cell. Using both experimentation and computer models we will delve deeper into how the emergent behaviour of these electrically excitable cells leads to the ability to control the heartbeat, kick a football, or think about why we are here!

Lec1 | Electrostatics & Debye screening: Nernst equation applied to intra-molecular interactions. The superposition principle; ionic bonds, hydrogen bonds, London dispersion forces can all be explained by Coulombic interactions; dipoles and dielectric effect; Bjerrum length calculation

- Discuss the consequences of the Nernst equation
- Discuss the extent to which life is an electrical phenomenon Lec2 | Introduction to neuroscience Membrane potentials, chemical synapses, gap junctions, calcium compartments
- Describe the structure and electrical signalling mechanisms of gap-junctions
  Lec4 | Hodgkin and Huxley: Hodgkin-Huxley model in neuroscience, solutions by simulations
- Simulate neuronal action potentials using Mathematica Lec5 | Optogenetics, behavioural feedback loops: Using light to control neural activity and probe the function of nervous systems
- Discuss, with examples, the utility of optogenetic manipulations in biology Lec6 | Control engineering: Resistors, capacitors, transistors, logic circuits
- Interpret and design logic circuits
  Lec7 | Ion Channels: structure and biological roles of ion channels 7 pumps

- Discuss and exemplify the classes, structures and biological roles of ion channels Lec8 | Neurons and neural networks: growth, insulation, synapsis of neurons
- Describe how neurons grow and synapse

#### B8 Synthetic biology

How can we use chemistry to make biological probes?

You will learn about protein synthesis and the key chemical reaction in this process, namely the formation of a peptide (amide) bond. But, you will use synthetic biology to design a new protein with a new amino-acid, a new codon, and a new translation apparatus. None of which exists in nature! Using the synthetic biology approach, you will learn about how normal protein synthesis is accomplished.

Lec1 | Reaction kinetics redux: the logic of chemistry

- Assemble rate equations for chemical reactions Lec2 | Molecular Modelling: Chemdraw. Pymol.
- Harness open source software to display and examine molecular structures Lec3 | Synthetic chemistry: Screening for activity
- Describe strategies for building, arraying and testing compound libraries Lec4 | Linkers: tagging, affinity techniques, crosslinking agents
- Exemplify the experimental uses of reversible and irreversible molecular tagging Lec5 | Solvation: phases, phase transitions
- Explain solvation mechanisms and their role in determining reactivity Lec6 | Ligand binding mechanisms
- Describe how metals bind and contribute to protein function Lec7 | Inhibitors and activators: Pharmacology
- Exemplify small molecule inhibition and activation of enzymes Lec8 | Genetic code expansion
- Discuss technological approaches to genetic code expansion

Labs: Over the 2 weeks of the block (6 lab sessions) students will use generic code expansion to engineer a protein incorporating a specific unnatural amino acid in a specific residue postion, express,

**B9** Light

Lec 1 - Ray optics 1

Overview view of light microscopy, Speed of light. Refraction, Refractive index, Focal points and principal planes, Lensmaker's equation.

Lec 2 - Ray optics 2

Image formation by a lens, Lateral and axial magnifications, System of lenses, Infinity optics Lec 3 - Wave optics 1

Maths for rotation, oscillation and wave, Light as electromagnetic wave, Dispersion, Polarisation Lec 4 - Wave optics 2

Complex exponentials, Interference/diffraction

Lec 5 - Light-matter interaction

Light as a quantum phenomenon, Fluorescence, LED, Image sensor

Lec 6 - Imaging by a microscope

Numerical aperture, Optical resolution, Fourier transform, Band-pass filters.

- Lab 1 Lenses, prisms, measurement of refractive index
- Lab 2 Collimation, diffraction by optical disks, Fraunhofer diffraction
- Lab 3 Measurement of PSF
- Lab 4 Photobleaching of GFP
- Lab 5- Stroboscopy 1 High-speed control of LED flashing
- Lab 6 -Stroboscopy 2 Measurement of the frequency of vibration by a smartphone
- Lab 7 Write paper
- Lab 8 Write paper
- B10 Pathogens and Parasites| Sam Dean

How have parasites evolved to invade our bodies?

Lec1 | Flagellar-driven motility

• Overview of eukaryotic flagella: their structure, how they are assembled and their varied functions

Lec2 | African Sleeping Sickness and trypanosomes

- Overview of Trypanosoma brucei: their lifecycle and how, despite being continuously exposed to host antibodies, they thrive in the host blood Lec3 | Intracellular parasites
- Overview of Leishmania and Trypanosoma cruzi, hugely successful kinetoplastid parasites that actually invade the cells that are supposed to kill them Lec4 | Gliding motility in apicomplexans
- How parasites move WITHOUT a flagellum and how is this machinery used to move through host cells and invade them
  - Lec5 | Plasmodium and Malaria

Plasmodium parasites that cause malaria, the big killer. How they resist the drugs that are supposed to kill them, and how humans have evolved to combat them Lec6 | Toxoplasma and toxoplasmosis

- Toxoplasma: "Why cats?" and how the parasite modifies the host behaviour (or not...) Lec7 | Co-evolution of parasites and the immune system
- How the evolution of pathogens and parasites has also driven the devlopment of our immune system and other host defences

## Learning outcomes

By the end of the module, students should be able to:

- Demonstrate the ability to apply creative analytical thinking in order to frame and answer scientific questions, especially about the structures, functions and mechanisms of cells and cellular organelles.
- Demonstrate a grasp of physical law as it applies to the structures and behaviours of living cells and their organelles, especially nuclei.
- Use mathematical approaches to solve problems relating to the structure and behaviours of living cells and their organelles.
- Describe and interpret quantitatively the mechanisms by which life harnesses electrical

forces

- · Access and use the scientific literature effectively
- Demonstrate the writing skills required to report experimental results in the format of a scientific paper, including the ability to write an abstract, to write a short critical review of the relevant literature, present results in an appropriate format and detail with appropriate statistics, discuss the results and frame a clear conclusion.
- · Harness computation to analyse scientific data
- Demonstrate the hands-on practical skills required to perform experimental tests of tractable scientific questions, especially about the structure and dynamics of cells and their organelles.
- Describe and interpret quantitatively how synthetic biology can be used to interrogate the mechanisms of life.
- Demonstrate a grasp of the properties of light and the principles of optical image formation
- Describe and discuss how pathogens and parasites have evolved to invade their hosts.
- Demonstrate the ability to accurately record experimental procedures and results in appropriate detail.
- Operate safely within a laboratory environment.

# Indicative reading list

**Biophysics and Physics** Physical Biology of the Cell Phillips et al Garland Science 2013 Molecular cell biology Molecular Biology of the Cell, 2014. Bruce Alberts et al ISBN: 9780815344322. **Biochemistry** Biochemistry (8th Edition) 2015. Jeremy Berg et al. ISBN-10: 1-4641-2610-0. Lewin's Genes XI, 2013. Krebs, Goldstein, Kilpatrick. ISBN: 128402721X Chemistry Principles of Modern Chemistry, 8th Edition 2016 Oxtoby, Gillis, Butler ISBN: 9781305079113 **Mathematics** Street-fighting mathematics 2010. Mahajan. ISBN : 978-0-262-51429-3 Measurements and their uncertainties 2010. Hughes & Hase. ISBN: 978-0-19-956632-7 Statistics at the bench 2010. Bremer & Doerge. ISBN: 978-0-87969-857-7 Basic mathematics for chemists Peter Tebbutt (2nd edition) 1998. Wiley ISBN: 978-0-471-97284-6 Computing Physical models of living systems Philip Nelson 2015 WH Freeman ISBN: 978-1-4641-4029-7 Programming in Python 3: A Complete Introduction to the Python Language (Developer's Library) Mark Summerfield ISBN: 978-0-321-68056-3 Math notes on Trigonometry, Basic Calculus, Complex Numbers, Taylor Series and Fourier Transform (https://moodle.warwick.ac.uk/course/view.php?id=38600)

# Interdisciplinary

Combines the methods of Biology, Chemistry, Physics and computing to interrogate the mechanisms of living systems.

# Subject specific skills

The ability to apply creative analytical thinking in order to frame incisive, tractable scientific questions, especially about the structures, functions and mechanisms of cells and cellular organelles.

The ability to demonstrate a grasp of physical law as it applies to the structures and behaviours of living cells and their organelles, especially nuclei.

The ability to solve problems relating to the structure and behaviours of living cells and their organelles.

Be able to describe and discuss how chemical biology and synthetic biology can be used to interrogate the mechanisms of life.

The ability to use mathematical and computational approaches to solve problems relating to biomolecular structures, functions and reactivities The ability to explain the organisation and behaviour of biomolecules, including the mechanisms by which biomolecules self-organise. The ability to design and describe simple optical circuits and to use focussed light to interrogate systems of biologically-relevant molecules.

# Transferable skills

Usage of mathematical and computational approaches to solve problems.

A grasp of safety rules and an ability to work safely in the laboratory environment.

The skills to accurately record experimental procedures and results, in appropriate detail.

The skills to use computational and statistical approaches to analyse data.

The skills to access and use the scientific literature effectively.

The writing skills required to report experimental results in the format of a scientific paper.

# Study

# Study time

| Туре              | Required                     |
|-------------------|------------------------------|
| Lectures          | 30 sessions of 1 hour (7%)   |
| Tutorials         | 30 sessions of 1 hour (7%)   |
| Practical classes | 30 sessions of 3 hours (20%) |
| Private study     | 150 hours (34%)              |
| Assessment        | 142 hours (32%)              |
| Total             | 442 hours                    |

Optional

10 sessions of 2 hours

# Private study description

Background reading in relation to each block (set of experiments), including reading scientific papers that are then referenced in the write ups.

# Costs

No further costs have been identified for this module.

## Assessment

You do not need to pass all assessment components to pass the module.

#### Assessment group D1

|  | Weighting  | Study time   | Eligible for self-<br>certification                          |  |
|--|--|--|--|--|
| Assessment component   |  |  |  |  |
| 3 x laboratory reports   | 40%  | 60 hours   | No   |  |
| Students work in pairs to ca<br>questions. At the end of eac<br>a report in the format of a so   | rry out laboratory expe<br>ch 2-week Block of the<br>cientific paper. Each rep | riments that addres<br>module, the results<br>port will be 1500 wc | s unsolved scientific<br>obtained are written up as<br>ords. |  |
| Reassessment component is the same   |  |  |  |  |
| Assessment component   |  |  |  |  |
| Mathematical Skills  | 10%  | 10 hours   | No   |  |
| Students will be tested on their mathematical skills from Block 6 with an in-class, short-answer test based on problem sets that they have been set during this block. |  |  |  |  |
| Reassessment component is the same   |  |  |  |  |
| Assessment component   |  |  |  |  |
|  |  |  |  |  |

No

| Assessing the proficiency in laboratory techniques, observing good laboratory practice, |
|---|
| engagement and contribution to group experiments.                                       |
|   |

Study time

Weighting

Reassessment component

Exemption

Under 'Principles and Scope' in the University's Examination and Assessment Policies, laboratory-based assessments are exempt from the right to remedy (1.3.2) as set out in Regulation 8-3 (iv) - "(if an appropriate standard in any required laboratory tests has not been achieved) candidates may be required to withdraw from their course".

Assessment component

Written examination 30%

Reassessment component is the same

Feedback on assessment

Laboratory reports - submission annotated and returned, general comments/what was good'/what could be improved' alongside marking rubric.

Assessment of laboratory skills - at the end of each two-week laboratory session, block leads will provide comments on proficiency, Good Laboratory Practice (GLP) and engagement/group contribution that arise. Due to the volume of feedback, stock phrases will be provided to the block leads, which may be amended or expanded at the lead's discretion. Further verbal feedback will be given to students on request.

In terms of practicalities, following GLP, executing the laboratory protocol, attaining proficiency in techniques taught and engaging/contributing to group activities (where required) will be based at 62 on the 20-point University scale. Exceptional attainment/contributions will grade higher, whereas disengagement, not observing GLP, and an unwillingness to acquire lab proficiency will score lower. Marks will not take into account whether a student achieved a desired experimental result or not. The block lead will work with the laboratory technician in observing and recording these across the cohort. Feedback will be provided biweekly at the end of each block.

Past exam papers for MD1A4

#### **Availability**

Courses

Eligible for selfcertification

No

No

This module is Core for:

• Year 1 of UMDA-CF10 Undergraduate Integrated Natural Sciences (MSci)