

2026 HIGHER DEGREE RESEARCH (HDR) INFORMATION BOOKLET

SCHOOL OF BIOTECHNOLOGY AND BIOMOLECULAR SCIENCES (BABS)

- GENOMICS AND BIOINFORMATICS
- MICROBIOLOGY AND MICROBIOMES
- MOLECULAR AND CELL BIOLOGY

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WELCOME FROM THE SCHOOL

This booklet provides a guide for students considering undertaking Higher Degree Research (HDR) in the School of Biotechnology and Biomolecular Sciences (BABS) at UNSW Sydney.

BABS is home to a number of active research groups that provide opportunities for postgraduate research students to work towards a PhD degree or a Masters by Research degree. For international students, we also offer a Master of Philosophy.

In BABS HDR programs candidates undertake a full-time research project supervised by a BABS researcher or approved external supervisor in an affiliated institution. HDR is immensely rewarding intellectually. All research in BABS is aimed at advancing science to make a real difference in the world. By investigating and understanding life at the molecular and cellular level, our students help solve real-world challenges.

Research in BABS is aligned to three discipline areas:

- » Genomics and Bioinformatics
- » Microbiology and Microbiomes
- » Molecular and Cell Biology

As you will see in this booklet, there is a wide scope of projects to interest candidates, with research spanning human bacterial pathogens, functional genetics, gene regulation, systems biology, viruses, cancer, neurobiology, extremophiles, synthetic and structural biology and more.

The work spans from hypothesis-driven 'blue sky' research that advances human knowledge, to application-focused research that has potential medical and industrial benefits for society.

Our HDR candidates benefit greatly from world-class facilities that include the Ramaciotti Centre for Genomics, which houses next-generation genomic sequencing technology.

Apart from imparting skills in scientific research, another aim of the BABS HDR programs is to equip students with skills in information technology, science communication and critical thinking, which will not only increase confidence but also make graduates more employable in an increasingly competitive workplace.

Our research community of staff and senior graduate students will do everything they can to ensure each candidate's experience is as enjoyable and scientifically stimulating as possible.

We invite you to become a part of our research effort by undertaking HDR with us.



Professor Shane Grey Head of School



Associate Professor Cecile King PGC - Admissions & Scholarships



Associate Professor Natalia Castaño-Rodríguez PGC - Candidature



Dr Michael Janitz PGC - Thesis & Examination

WHY DO A PHD IN BABS?

The Doctoral degree programs within the School of Biotechnology and Biomolecular Sciences (BABS) provide the highest level of training in key areas of scientific research. Students work on an independent research project encompassing the broader interests of one of the research teams within the School. In the early stages of the program, students receive close supervision and guidance in the management of their project. In the later stages, however, students are encouraged to exercise initiative and demonstrate originality. In the last year of the program, the candidate should be able to work independently and be guided rather than directed by the supervisor.

A key benefit of doing a PhD in BABS is that it provides an active, hands-on learning experience in a scientific research environment. Students become part of a research team within a lab in the School, with supervisory oversight provided on an individual basis by an experienced academic. In addition, interaction with other experienced researchers within the group in an informal, relaxed atmosphere complements the formal part of the PhD program, of completing the predetermined research project and writing a thesis.

The Program is designed to provide advanced training and knowledge in one of the School's majors:

- » Biotechnology
- » Genetics
- » Microbiology
- » Molecular and Cell Biology

A PhD is also an opportunity for the student to reflect on their future career.

PhD graduates have the opportunity to develop greater competence and confidence in the practical skills and laboratory methods acquired during their Honours year, while developing key attributes sought by employers, including:

- » Development of critical thinking skills
- » Extensive use of a variety of information and communication technologies
- » Familiarity with a range of computer software for oral and written presentations
- » Training in online database manipulation and data analysis
- » Collaboration in industrial research and commercialisation of science nationally and internationally

The higher level of such attributes are well recognised by employers and greatly increase the possibility of gaining employment in industry, agriculture, medical or research organisations.

A PhD degree requires three to four years full-time study and completion of a written thesis. The length of a doctoral thesis is normally around 100,000 words. The thesis is reviewed by members of the Australian and international scientific academic community. In the course of their research, PhD students must make a distinct contribution to the knowledge within their specific discipline. Ideally, this will result in the publication of original research findings in peer-reviewed journals of international standing.

Who is eligible for PhD?

The minimum entry requirement for a PhD is a 4-year Bachelor's degree with First or Upper Second Class honours; or with the consent of the potential supervisor, a qualification or combination of qualifications considered to be equivalent by the Faculty of Science Higher Degree Committee e.g. completion of a Bachelor degree and substantial laboratory experience.

It is essential for you to have identified a BABS supervisor who has agreed to take you on as a student prior to applying.

Applicants must provide evidence that their English language ability meets the minimum requirements for admission: English language requirements.

Further information can be obtained from the Graduate Research School (GRS).

PhD Program Options

PhD - Biochemistry & Molecular Genetics (1410)

PhD - Bioinformatics (1683)

PhD - Biotechnology (1036)

PhD - Microbiology & Immunology (1440)

FIVE REASONS STUDENTS CHOOSE TO DO A PHD

Biotechnology and Biomolecular Sciences are rapidly growing areas with broad impact. There are many rewarding career paths that require advanced knowledge in life sciences and many reasons that people decide to undertake a PhD within the School of Biotechnology and Biomolecular Sciences. Here's just a few:

Salary

Money can't buy happiness, but it doesn't hurt. UNSW students are among the highest earning undergraduates in Australia [1] and our post-graduate students are the highest earning in the country [2]. The median salary for a UNSW undergraduate student is \$77,500 and with post-graduate studies, UNSW students' salary increases to \$93,600. A PhD can also open avenues to more opportunities and better career progression.

Employability

UNSW post-graduates have an impressive track record of stepping out of post-graduate degrees and into full-time work. From 2018-2020, 82.5% of UNSW post-graduates research students stepped into full-time employment. Overall students with post-graduate degrees maintained a high employment rate during COVID-19 economic downturn of 2020. A PhD in BABS can help future-proof your career.

Make A Difference

Understanding and harnessing the information encoded within a cell is one of the great challenges of the 21st century. We've already seen how this can change lives (think personalised genomic medicine, CAR-T cells, and CRISPR). Research in BABS seeks to understand diverse aspects of molecular and cellular biology and translate that knowledge into real world solutions. PhD projects in BABS lead to advances in genetics, neurobiology, metabolism, infectious diseases, and environmental sciences. A PhD in BABS gives you to tools to make a difference.

Find The Cutting-edge And Stay There

Technology is advancing at an unprecedented pace and high-quality jobs require advanced, specialist knowledge. A PhD in BABS places you at the forefront of the latest advances in life sciences and equips you with the skills to stay there.

Biotechnology And Biomolecular Sciences Are Growth Areas

Investment in biotechnology has grown rapidly for the past decades and continued to grow strongly in 2020 [3]. Investment in biomedical research returns both financial, health, and productivity gains [4] and perhaps unsurprisingly biomedical research in Australia continues to be supported by both the Government and private sector. Several new medical research institutes have recently been created - including the Randwick Health and Innovation Precinct across the road from BABS. Biotechnology and Biomolecular Sciences are areas of strong growth that can provide rewarding and lasting career paths.

Lastly, you might want to read this recent paper on 'Research Culture: Highlighting the positive aspects of being a PhD student'.

- 1. QILT, Nov 2020, <u>2020 GOS National Report Graduate Outcomes Survey: Short-term graduate</u>
 Outcomes in Australia.
- 2. Menzies, S, Jan 2019. <u>UNSW among top Australian universities for high salaries</u>.
- 3. DeFrancesco, L. Financing breaks all records in 2020. *Nature Biotechnology*. 2021. 39, p133.
- 4. Association of Australian Medical Research Institutes. Oct 2018. Economic impact of Medical Research in Australia.

OTHER HDR OPPORTUNITIES IN BABS

Masters by Research (MSc)

Each Masters program is an advanced area of study where graduates may obtain specialist knowledge in a particular area of science. These research programs focus on training students to be innovative and independent.

The MSc degree is normally two years full-time in duration and students are required to dedicate most of their time to research and the preparation of a Masters thesis. The length of a Masters research thesis normally should not exceed 75,000 words. Once completed, the thesis is examined by members of the Australian and international scientific academic community.

Who is eligible for MSc?

The minimum requirement for admission is a relevant 4-year Bachelor's degree with Honours that includes a substantial research component; or with the consent of the potential supervisor, a qualification or combination of qualifications considered to be equivalent by the Faculty of Science Higher Degree Committee.

It is essential for you to have identified a BABS supervisor who has agreed to take you on as a student prior to applying.

Applicants must provide evidence that their English language ability meets the minimum requirements for admission: English language requirements.

Further information can be obtained from the Graduate Research School (GRS).

MSc Program Options

MSc - Biochemistry & Molecular Genetics (2460)

MSc - Biotechnology (2036)

MSc - Microbiology & Immunology (2490)

Master of Philosophy (MPhil)

The Master of Philosophy (BABS) is offered to international students and is recognised as a postgraduate research degree that sits somewhere between a BSc and a PhD. It is designed to be completed over six terms (full-time), or 1.5 years, during which three subjects of coursework are undertaken and a supervised research project is completed in the supervisor's research laboratory.

The outcomes of the research project are documented in a thesis, which is examined.

This program is intended for international students who do not meet the requirements for an MSc by Research or PhD, or those wishing to develop expertise in an area different from their undergraduate degree. The MPhil should be considered as an alternative to an Honours year for international students.

This qualification allows students to experience modern and sophisticated laboratory techniques that apply to a wide range of biotechnology and molecular biology fields. To allow greater flexibility to pursue a particular area of interest, course electives may be chosen, subject to the approval of the PGC.

Who is eligible for MPhil?

International students with a first class degree or 4-year degree in a relevant discipline.

It is essential for you to have identified a BABS supervisor who has agreed to take you on as a student prior to applying.

Applicants must provide evidence that their English language ability meets the minimum requirements for admission: English language requirements.

Further information can be obtained from the <u>Graduate Research School (GRS)</u>.

MPhil Program

MPhil - Science (2475)

UNSW SCHOLARSHIPS

UNSW Sydney Raises Stipends for Higher Degree Research Scholarships

International Research Scholarships

UNSW Sydney offers a number of prestigious scholarships to International Higher Degree Researcher including:

- · Australian Government Research Training Program (RTP) Scholarship
- University International Postgraduate Award (UIPA)
- · Tuition Fee Scholarship (TFS) plus a Research Stipend

Further information including how to apply can be obtained from the Graduate Research School (GRS).

Domestic Research Scholarships

UNSW Sydney assists Domestic Higher Degree Researchers through a range of scholarship schemes including:

- Australian Government Research Training Program (RTP) Scholarship
- University Postgraduate Award (UPA)
- Aboriginal and Torres Strait Islander Research Training Program Scholarships
- Faculty Top Up Scholarships attached to RTP Scholarship or UPA

Further information including how to apply can be obtained from the **Graduate Research School (GRS)**.

UNSW / Home Country Joint Scholarships

A number of UNSW / Home Country Joint Scholarships have been established to promote international collaboration for international candidates seeking to undertake PhD degree at UNSW including:

- China Scholarship Council (CSC) Scholarship
- UNSW / HEC Pakistan Joint Scholarship
- UNSW / ANID Chile Joint Scholarship
- · Vietnam Vingroup Scholarships
- GOstralia! Higher Degree Research Scholarships

Further information including how to apply can be obtained from the **Graduate Research School (GRS)**.

Other Scholarship Opportunities

- Faculty and Donor Funded Scholarships Further information including the current range of full or top up scholarships available to candidates can be obtained from the Graduate Research School (GRS).
- Externally Funded Scholarships Further information including eligibility criteria, funding availability and application processes can be obtained from the <u>Graduate Research School (GRS)</u>.

Additional Information

UNSW SCHOLARSHIPS

HDR Development and Research Training Grant (DRTG)

The DRTG scheme is a centrally funded Grant-in-Aid program that provides 2nd and 3rd year candidates with an opportunity to enhance their candidature experience by providing funding to attend conferences, as well as profile building and skill development activities.

Throughout an individual's candidature, they can apply twice to the DRTG scheme, where the maximum funding being requested is \$1,500 per round (\$3,000 in total over the two rounds). This funding is a contribution to the total cost of the activities, and may not necessarily fund these activities outright.

Candidates can request funding to support one or more of the five listed activities:

- · Registration fees for a virtual conference (international or domestic)
- Registration, travel and/or accommodation fees for a domestic conference
- · Registration, travel and/or accommodation fees for international conference
- Professional Development short courses and training opportunities
- Cost of childcare where it allows the candidate to attend one of above approved activity types

Note: This scheme was on offer in 2025. The 2026 scheme is to be confirmed. Further information is available on the <u>UNSW HDR Hub</u> which can only be accessed after enrolment is completed (it is only available to current students).

UNSW PhD Candidate Paid Parental Leave Scholarship

The PhD Candidate Parental Leave Scholarship is open to PhD candidates who have been awarded a scholarship with no parental leave entitlements.

Candidates successful for this scholarship receive paid parental leave for up to 12 weeks equal to the value of their scholarship.

Further information is available on the <u>UNSW HDR Hub</u> which can only be accessed after enrolment is completed (it is only available to current students).

UNSW Science PhD Scholarships

The Faculty of Science offers a Science PhD Maternity Scholarship for women PhD research students who suspend their enrolment for a session to have a child. They aim to support women in research by bridging the gap and offering financial support during maternity leave.

The Faculty also offers the Prof. Joseph P Forgas PhD Travel Award. The purpose of this award is to provide an international learning, research, and networking opportunity for PhD candidates studying in the Faculty of Science. The Award will support travel and associated costs to attend an international university in aid of research progress, to promote collaborations and develop the candidate's professional career. Travel will preferably be for a minimum of 3 months with preference given to students travelling to universities in Europe and the USA.

Further information including eligibility requirements and how to apply can be obtained from <u>UNSW Science</u> - <u>PhD Maternity Scholarship</u> and <u>PhD</u> scholarships.

BABS SCHOLARSHIPS

The Adrian Lee Travel Scholarship

This Scholarship is a School-funded initiative to provide eligible postgraduate research candidates with the opportunity to undertake study, learn new techniques, collaborate or explore opportunities for collaboration with other labs, universities or research institutions.

The value of the scholarship is \$6,000 maximum, payable in one lump sum. The Scholarship is tenable for one year only (travel must be taken in the year of award).

The scholarship can be used for travel, reagents or project costs, accommodation and other related travel costs.

Once enrolled, candidates can access the full guidelines including eligibility requirements and how to apply via the BABS HDR SharePoint.

BABS HDR Development & Conference Grant

This scholarship is a school-funded initiative to provide eligible PhD & MSc research candidates with the opportunity to:

- Present their research at conferences (domestic and international) and/or
- Complete professional development short courses and training opportunities if required and appropriate

Eligible candidates can access funding up to \$3,000 maximum throughout their degree, with \$2,000 being the maximum award for international travel and \$1,000 being the maximum award for domestic travel.

Once enrolled, candidates can access the full guidelines including eligibility requirements and how to apply via the BABS HDR SharePoint.

HOW TO APPLY

Ready to apply for a research higher degree program at UNSW? Below outlines the steps you need to follow, including the supporting documents you need to provide, and the key dates for your application.

It is the applicant's responsibility to ensure that their application is submitted in full by the scholarship closing date. All correct and satisfactory documents must be fully submitted with the application, including English translations, Financial Declaration and proof of English. Additional documents can't be added to an application until after the application has been reviewed, which may not occur before application deadlines. Please refer to Step 3 to ensure you have the correct documents before submission.

Step 1: Determine Your Eligibility

There are a number of eligibility requirements you need to ensure you meet prior to applying for a Higher Research Degree. See here.

Step 2: Find a Supervisor and Prepare Your Research Description

Finding a supervisor with compatible research interests and working styles is critical to your success as a HDR candidate. Information on HDR supervisors and research areas can be found in this booklet or on the BABS website.

Once you have decided which supervisor you wish to contact for further discussion, email is the preferred method of contact. It is essential to spend some time with prospective supervisors to discuss the details of available projects before submitting your application. In your email, please ensure that you:

- a. Identify which research area you are interested in, and why
- b. Indicate which term you intend on commencing (Term 1, 2 or 3)
- c. Advise your availability times for an interview
- d. Attach a copy of your CV and academic transcript
- e. Confirm you have available funding to cover both living expenses and the tuition fees
- f. Indicate if you have appropriate visa status

More advice and resources for finding a supervisor is available here.

Step 3: Prepare Your Supporting Documentation

There are a number of supporting documents that you are required to submit with your application. Full details are available here.

Step 4: English Translations of Documents

Full details are available here.

Step 5: Meet the UNSW English Language Requirements

Full details are available here.

Step 6: Submit an Application

Once you have secured a supervisor, held your interview, developed a research description, and prepared your supporting documents, you are ready to Iodge your application here. Only full applications (i.e. with all required documents) will be processed for assessment.

If you wish to be considered for a scholarship, simply indicate this on the application form and select the applicable scholarship round for your preferred term start date. For more information on applying for a scholarship, please visit Graduate Research Scholarships.

BABS program codes can be found in this booklet or on the <u>BABS website</u>. You can confirm the correct program code with your supervisor.

Application deadlines can be found online at Key Dates.

Further details about after you submit an application, receiving an offer and responding to an offer are available here.

BABS HDR inquiries

BABS HDR Support Officer, BABS School Office Room 520, Biological Sciences Building D26 T+61 (0)2 9065 9717

E babs-pg@unsw.edu.au

RESEARCH SUPERVISORS -

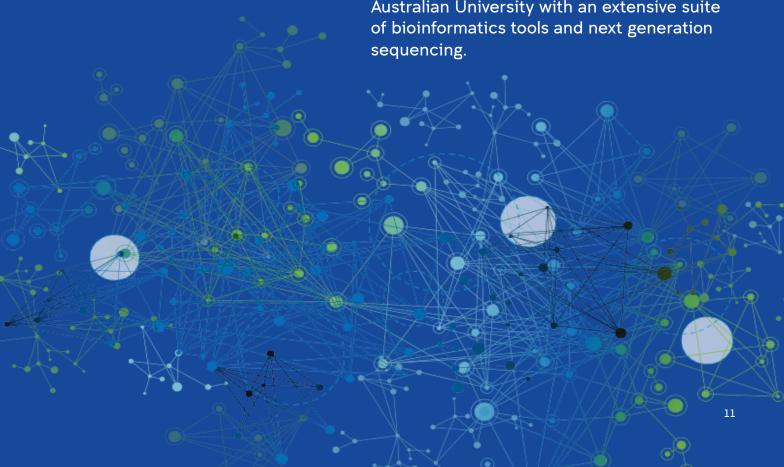
GENOMICS AND BIOINFORMATICS

CLUSTER STRENGTHS:

- » Gene Regulation
- » Systems Biology
- » Neuogenomics

Genomics and Bioinformatics is an invaluable hybrid of science, concerning the structure and function of genomes and the use of computational technology to capture and interpret biological data. While scientists previously focused on singular cells, the enormous development in bioinformatics over the last decade has enabled us to study cells on a mass scale.

We are focused on enabling medical breakthroughs and clinical application with our access to cutting- edge computational biology. UNSW Biotechnology and Biomedical Sciences houses the Ramaciotti Centre for Genomics, the largest and most comprehensive genomics facility at any Australian University with an extensive suite of bioinformatics tools and next generation sequencing.





merlin-crossley

W research.unsw.edu.au/people/professor-



Transcription factors and gene regulation in blood cells.



We study how transcription factors control cell fate and how the breakdown of this process leads to disease. We apply this knowledge with the ultimate aim of developing the next generation of artificial transcription factors and to develop new therapeutic strategies for blood diseases. Currently, our collaborative research group includes a lab manager, 7 PhD students and 2 Honours students.

RESEARCH GOALS

- · Fundamental mechanisms of gene regulation
- Epigenetics
- Artificial DNA-binding proteins
- Blood development
- Haemophilia
- Sickle cell anaemia and thalassaemia

RESEARCH IN DETAIL

We are interested in understanding the fundamental molecular mechanisms by which genes are controlled. Ultimately we hope to treat chronic diseases, such as sickle cell anaemia, by turning on beneficial genes or turning off harmful genes.

BIOGRAPHY

Merlin Crossley is Deputy Vice-Chancellor (Academic and Student Life) at UNSW and Professor of Molecular Biology. He has also worked or studied at the Universities of Melbourne, Oxford, Harvard and Sydney. He has been recognized by numerous awards, including a Rhodes Scholarship, the Australian Academy of Science's Gottschalk Medal and the Australian Society for Biochemistry and Molecular Biology's Lemberg Medal. He has made significant contributions to academic administration, serving as Dean at UNSW, and Acting Deputy Vice-Chancellor Research at the University of Sydney.

RECENT PUBLICATIONS

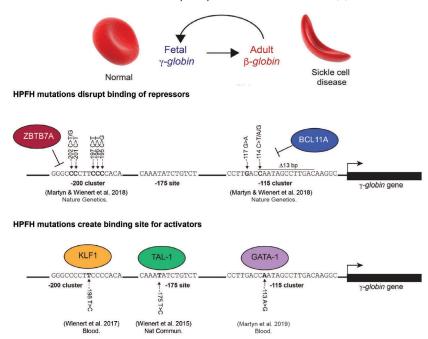
'Disrupting the adult globin promoter alleviates promoter competition and reactivates fetal globin gene expression.' *Blood*, 2022, 139(14):2107-2118

'Methylation of a CGATA Element Inhibits Binding and Regulation by GATA-1', *Nature Communications*, 2020, 11(1):2560

'Natural regulatory mutations elevate the fetal globin gene via disruption of BCL11A or ZBTB7A binding.' *Nature Genetics*, 2018 50(4):498-503

'Transcription factors LRF and BCL11A independently repress expression of fetal hemoglobin', *Science*, 2016, 351(6270):285-9

'Directing an artificial zinc finger protein to new targets by fusion to a non-DNA-binding domain', *Nucleic Acids Research*, 2016, 44(7):3118-30





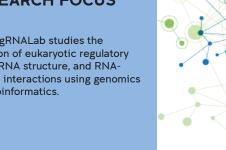
Dr Selene Fernandez **ARC Future Fellow**

Room 301D, Level 3 Bioscience South Building E26 E s.fernandez_valverde@unsw.edu.au W regrnalab.github.io

W unsw.edu.au/staff/selene-fernandez-valverde



The RegRNALab studies the evolution of eukaryotic regulatory RNAs, RNA structure, and RNAprotein interactions using genomics and bioinformatics.



RESEARCH PROGRAM

Most of our genes are largely shared with worms, flies, and even marine sponges - how do we look so different? How can we consistently go from an embryo or a seed and form a full human or plant in a precisely controlled manner? How can loss of this control can lead to disease? In the RegRNALab, we leverage bioinformatic and genomic data analysis to understand how gene regulatory mechanisms change over time while maintaining the intricate development and diversity of cells found in multicellular plants and animals. We are interested in the role regulatory RNA molecules play in these processes, in particular non-coding RNAs. To tackle these questions, we study several biological models that span different evolutionary scales, including humans, plants, and animals.

RESEARCH PROJECTS

1. Identification of long non-coding RNAs (IncRNAs) under recent selection in human populations (computational).

In collaboration with the group of Dr. Andres Moreno-Estrada we recently characterized the genetic variation of modern Mexicans across Mexico. We are leveraging this and other Biobanks to identify IncRNAs under selection and associated with cardiometabolic syndromes in admixed populations. We are very interested in how genetic diversity and admixture in humans impact disease.

2. Identification and characterization of non-coding RNAs in atherosclerosis (computational/experimental)

In collaboration with Dr Ashish Misra, we are working on identifying IncRNAs relevant to atherosclerosis. Atherosclerosis is the leading cause of death globally and in Australia, where it is responsible for 1 in 4 deaths. Despite such a high disease burden, available therapies have low efficacy; hence, finding novel molecular targets and biomarkers, such as IncRNAs, can pave the way for novel therapies and diagnostic methods.

3. Identification of tissue and cell type-specific RNA motifs (computational)

One of the greatest challenges in RNA therapy development is the limited specificity of delivery vehicles, such as lipid nanoparticles. We are looking to computationally identify novel signatures in RNAs specific to tissues and cell types to enable specific transcription that only occurs in targeted cells.

4. Characterization of binding sites & structures of RBP-bound RNAs in plants (experimental/ computational)

As part of an ARC Future Fellowship awarded to Dr Fernandez, we are characterizing the RNA bound by multiple RNA Binding Proteins (RBPs) and the structures they form to find general rules that can be used to predict binding using machine learning methods.

BRIEF BIO

Dr Fernandez is a Senior Lecturer funded by the UNSW RNA Institute. She holds a BSc in Genomic Sciences from the National Autonomous University of Mexico and a PhD and postdoc from The University of Queensland. She was a principal investigator at the Advanced Genomics Unit, Cinvestav, from 2016 to 2023, before joining UNSW as a Senior Lecturer in mid-2023. Dr Fernandez was awarded the Mexico national scholarship L'Oreal-UNESCO-AMC-CONACYT for Women in Science in 2016, the International Rising Talents Fellowship of L'Oreal-UNESCO in 2018, and the Gender Equality Award by the International Union of Immunological Societies in 2019. She currently serves on the board of the International Society for Computational Biology (ISCB) and is an editor for the journals PLoS Biology and Genome Biology and Evolution.

You can see the most recent lab publications here.



Dr Michael Janitz Senior Lecturer Room 3106, Level 3 West Bioscience South Building E26 E m.janitz@unsw.edu.au W babs.unsw.edu.au/michael-janitz



Human transcriptome, circular RNAs, neurological disorders and cancer, biomarkers of complex diseases.



SCIENTIFIC INTEREST

Michael is an internationally recognised expert in the field of transcriptomics, next-generation sequencing technologies and bioinformatics. His current scientific interest focuses on investigations of gene expression and alternative splicing patterns in distinct structures and cell types of the healthy human brain and perturbation of transcriptome profiles during the onset and progression of neurological disorders including multiple system atrophy, epilepsy, amyotrophic lateral sclerosis and chronic fatigue syndrome. Michael is also interested in exploration of cancerspecific alterations in transcriptome profiles towards identification of novel biomarkers such as circular and non-protein coding RNAs.

RESEARCH CONTRIBUTIONS

During his PhD studies and subsequently as a postdoc at the German Centre for Rheumatism Research, Michael specialised in investigating the influence of the sequence polymorphism within the promoter regions of MHC class II genes in several inbred mice strains (Janitz et al. 1997; Janitz et al. 1998; Cowell et al. 1998).

Joining the Max Planck Institute for Molecular Genetics (MPIMG) converged with Michael's growing interest in studying transcription at the genome-wide level. Amongst others, he was involved in collaborative projects to characterise cDNA sequences on the level of the whole transcriptome in mice T helper cells and bovine brain (Gutjahr et al. 2005; Jann et al. 2006), respectively.

While at the MPIMG, Michael and his research group focused on developing a transfected-cell array as a high-throughput genomic tool for functional analysis of genes and their products (Vanhecke and Janitz 2004). This resulted in application of the cell arrays for subcellular protein localisation studies (Hu et al. 2006; Hu et al. 2009; Hu et al. 2010), protein-protein interaction screens (Fiebitz et al. 2008), and functional promoter analysis (Cheng et al. 2010). In addition, with collaborative partners in national and European Community research programs, he applied gene

expression profiling studies to identify the genes involved in T helper lymphocytes type 1 immune response (Niesner et al. 2008) and differentiation of murine palatal development (Nogai et al. 2008). His research group also developed miniaturised microarray platforms for DNA hybridization studies using PNA-(Bauer et al. 2004) and LNA-modified oligonucleotide probes (Guerasimova et al. 2006; Liu et al. 2006 and 2007), thus contributing to more efficient exploration of the genome structure and function.

CURRENT RESEARCH

Since his appointment at UNSW Michael has been focused on exploration of different segments of human and mouse transcriptome using combination of Illumina and nanopore DNA and RNA sequencing as well as unique analytical pipeline combining in-house and publicly available bioinformatics tools. He and his team provided first insights into non-protein coding transcriptome of different regions of the human cortex, both in health (Mills et al. 2013; Mills et al. 2015a; Mills et al. 2015b; Bliim et al. 2019) and neurological diseases such as Alzheimer's disease (Twine et al. 2011; Mills et al. 2013; Mills et al. 2014), multiple system atrophy (MSA)(Mills et al. 2015c; Mills et al. 2016) and epilepsy (Mills et al. 2020) as well as cancer (Takenaka et al. 2016; Chen et al. 2017a). His recent research has been concentrated on circular RNAs (circRNAs) genome-wide expression patterns in the normal human (Chen et al. 2019) and mouse brain (Chen et al. 2018) as well as MSA (Chen et al. 2016) and endometrial cancer (Chen et al. 2017b).

Current research projects aim at understanding of circRNA-miRNA-mRNA networks in epilepsy and amyotrophic lateral sclerosis as well as gynaecological malignancies. Another emerging avenue of Michael's research is exploration of RNA post-transcriptional modifications in healthy human tissues and cancer using nanopore direct RNA sequencing. Michael's quest for the discovery of circRNAs, which might serve as molecular biomarker for MSA and endometrial cancer, constitutes another important element of his research program.

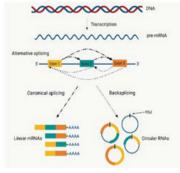


Figure 1. The formation of linear mRNAs and circular RNAs through canonical splicing and backsplicing, respectively. The mechanism of backsplicing leads to covalent linkage of the downstream 3'-end of a pre-mRNA sequence to an upstream 5'-end of the pre-mRNA strand. This process leads to generation of a backspliced junction (BSJ), denoted by the black line in circular isoforms, whoch is a unique feature of circRNAs. Linear mRNAs are formed through the canonical splicing process whereby introns are excised from the pre-mRNA strand, forming exonic isoforms of linear mRNA with no BSJ (adapted from Curry-Hyde et al. 2020).



Associate Professor Cecile King Room 3108, Level 3 East Bioscience South Building E26 E c.king@unsw.edu.au W babs.unsw.edu.au/cecile-king

RESEARCH FOCUS

RNA at the immune interface; regulation of the immune response by RNA, transposable elements, RNA sensing and type-1 interferons in immune responses and autoimmune disease.

RESEARCH PROGRAM

One of the most intriguing aspects of the immune system is its ability to react to foreign invaders, but not to self. To distinguish healthy cells (self) from infected cells or pathogens, the immune system utilizes multiple receptors and signalling pathways that enable specific recognition of proteins and nucleic acids. Recognition of foreign RNA occurs inside cells and depends upon unique features of pathogen RNA that allow our RNA sensors to see it as foreign. However, for some people, the immune system reacts inappropriately to self-RNA and this can trigger autoimmunity.

The achievement of self-tolerance is even more impressive when we consider that we are colonized by multiple microbial species that are important for our health, and our genomes harbour an abundance of retrovirally sourced nucleic acid acquired from past encounters with retroviruses, known as retroelements. Whilst a challenge for self-tolerance, the retention of retroelements in the genome has been shown to serve multiple regulatory functions that influence the transcription of protein coding genes. Our own work has shown that retroelements can function to neutralise the deleterious effects of duplicated genes in immune gene families, favouring host survival during virus infection.

We are employing both an inside and outside approach to study RNA in the immune system-by studying how the immune system senses RNA during immune responses and how retroelement RNA's regulate immune responses.

Project 1: The role of retroelements in regulation of immune responses. This project will take a genomewide approach to analyses the role of retroelements in the immune system and their relationship with immune genes families that have arisen by gene duplication. The project will be jointly supervised by Professor Marcel Dinger (babs.unsw.edu.au/marcel-dinger).

Project 2: RNA sensing in health and disease. This project will probe the underlying mechanisms of RNA sensing by the immune system and how autoimmune disease associated mutations in RNA recognition pathways influence immune responses.

More detailed information on specific projects and ongoing research is available at: babs.unsw.edu.au/cecile-king

BIOGRAPHY

Associate Professor Cecile King joined the School of Biotechnology and Biomolecular Sciences at the University of New South Wales Sydney in 2021. Cecile received her Ph.D. in Immunology from the University of Western Australia and completed her postdoctoral training at the Scripps Research Institute, La Jolla, USA. She joined the faculty of the Garvan Institute for Medical Research, Sydney, Australia in 2005, where she established her independent research program. Cecile's research has made important contributions to our understanding of T cell subsets and cytokines in adaptive immune responses and autoimmunity. Cecile's current research focus is RNA at the immune interface: Understanding both immune recognition of RNA and immune regulation by RNA.

Contact Cecile about research supervision opportunities: c.king@unsw.edu.au



Professor John Mattick Room 3111, Level 3 East Bioscience South Building E26 E j.mattick@unsw.edu.au W babs.unsw.edu.au/john-mattick

RESEARCH FOCUS

regulating human development and cognition

The role of noncoding RNA in

The human genome contains just 20,000 proteincoding genes, similar in number and with largely the same functions as those in other animals, including tiny nematode worms that have only 1,000 cells. By contrast, the extent of non-protein-coding DNA increases with developmental complexity, reaching 98.7% in humans. Moreover, these noncoding sequences, previously thought to be junk, are differentially transcribed to produce hundreds of thousands of small and long non-protein-coding RNAs that are expressed intronically, intergenically and antisense to protein-coding genes.

Most long noncoding RNAs exhibit highly specific expression patterns and subcellular locations, and appear to be expressed from genetic loci termed enhancers that control the spatiotemporal patterns of differentiation and development. Many have evolved rapidly under positive selection for adaptive radiation, and increasing numbers have been shown to have key roles in development, brain function, cancer and other disorders.

RESEARCH AREAS

- The structure-function relationships in regulatory RNAs
- The roles of long noncoding RNAs in human development and cognition
- The roles of noncoding RNAs in complex diseases

PROJECTS

- Bioinformatic and functional analyses of long noncoding RNAs expressed from genomic regions associated with human intelligence, mood disorders, addiction, personality and other neurological traits.
- Bioinformatic and functional analyses of long noncoding RNAs expressed from human genomic regions associated with immunological traits and disorders, including autoimmune
- Bioinformatic analysis of the structure-function relationships in long noncoding RNAs.

RECENT PUBLICATIONS

- Mattick JS, et al. (2023) Long noncoding RNAs: Definitions, functions, challenges and recommendations. Nature Reviews Molecular Cell Biology 24: 430-447. https://doi.org/10.1038/ s41580-022-00566-8.
- Mattick JS (2023) A Kuhnian revolution in molecular biology: Most genes in complex organisms express regulatory RNAs. BioEssays 45: article 2300080. https://doi.org/10.1002/bies.202300080.
- Mattick JS & Amaral PP (2022) RNA, the Epicenter of Genetic Information. CRC Press (free online). https://doi.org/10.1201/9781003109242.

BIOGRAPHY

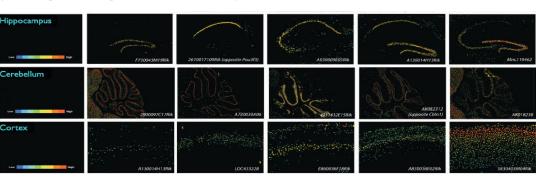
Professor Mattick is the Professor of RNA Biology. He was previously the Director of the Institute for Molecular Bioscience at the University of Queensland, the Australian Genome Research Facility and the Garvan Institute of Medical Research. He is a Fellow of the Australian Academies of Science, Technology & Engineering and Health & Medical Sciences.

Professor Mattick is recognised for having pioneered a new understanding of the information content and function of the genome in other developmentally complex organisms. He has published over 300 research articles and reviews, which have been cited over 100,000 times.

Professor Mattick's honours include Membership of the European Molecular Biology Organisation, Honorary Fellowship of the Royal College of Pathologists of Australasia, the Australian Society for Biochemistry and Molecular Biology Lemberg Medal, the International Union of Biochemistry and Molecular Biology Medal, the University of Texas MD Anderson Cancer Center Bertner Award for Distinguished Contributions to Cancer Research and the Human Genome Organisation Chen Medal for Distinguished Achievement in Human Genetics and Genomic Research.

Contact John about research supervision opportunities: j.mattick@unsw.edu.au

Figure: Images of the restricted expression of noncoding RNAs in the brain.





Dr Emily Oates Senior Lecturer Room 320C, Level 3 Biological Sciences North Building D26 E e.oates@unsw.edu.au W babs.unsw.edu.au/emily-oates

Human neuromuscular disease gene discovery, the clinical and genetic characterisation of new and emerging neuromuscular disorders, disease-related pharmacogenomics, muscle transcriptomics, clinical severity prediction, genetic treatment development.

Our main area of research interest is the discovery of new genes responsible for congenital and childhoodhood onset genetic muscle conditions. This group of disorders causes significant physical disability and sometimes early death.

Around half of all individuals with one of these conditions still do not have a genetic diagnosis. This adversely impacts their clinical care and the care of their family. In many cases a genetic diagnosis has not been established because the causative gene is not yet known to be a disease-causing gene.

Another significant challenge is that there are no available treatments to prevent, halt, or slow the progression of most congenital and childhood-onset muscle diseases - even when the genetic basis of the disease is known.

Our research is focused on the discovery of new childhood-onset neuromuscular disease genes, and the analysis of the functional and clinical impacts of disease-causing variants within these genes using state-of-the-art phenotyping, cell and tissue-based analyses, DNA and RNA sequencing and bioinformatic approaches. We use this information to increase genetic diagnosis rates for affected individuals and their families, and to advance our understanding of the clinical characteristics, natural history, and underlying pathogenesis of these disorders. We also have emerging interests in disease-related pharmacogenomics, muscle transcriptomics, clinical severity prediction and the development of new genetic treatments using insights gained through the detailed analysis of the genetic basis of muscle disorders.

BIOGRAPHY

Dr Emily Oates is a head of the UNSW Medical Genomics Group, a neurogenetics consultant for the Sydney Children's Hospital Network, and senior lecturer (UK), The Broad Institute (USA) and NIH (USA). in medical genomics. Following paediatric physician training she completed specialist clinical genetics training, a neurogenetics-focused PhD and postdoctoral studies in both Australia and London, UK (Great Ormond Street Hopital). She has 14 years of clinical experience in the diagnosis and management of infants and children with neuromuscular disorders. She also has extensive expertise in the clinical characterisation of new neuromuscular disorders and the analysis of human genomic data for diagnostic and gene discovery purposes. She sees patients weekly in the Sydney Children's Hospital Neuromuscular clinic. Many of her patients are enrolled in her research projects. Dr Oates recently completed an NHMRC Neil Hamilton Fairley Early Career Fellowship. This fellowship was focused on harnessing state-of-the-art massively parallel DNA and RNA sequencing technologies to improve genetic

diagnosis rates for patients with neuromuscular disorders and to identify new disease-causing genes. During her PhD and ECR fellowship she contributed to the identification of multiple new disorders including BICD2-SMA and SCN4-myopathy. She also authored multiple peer reviewed journal articles, including a large international publication aimed at comprehensively characterising the genetic and clinical features of the largest congenital (recessive) titinopathy cohort ever described at that time (84 authors: top 10 most downloaded 2018-2019 Annals of Neurology papers). Recessive titinopathy is a significant ongoing clinical and research interest for Dr Oates and her team.

Dr Oates is also lead investigator for a new (2022) Medical Research Future Fund (MRFF) grant: 'Advancing congenital and childhood-onset muscle disease diagnosis and treatment - a cross- disciplinary Australian collaboration'. This grant will fund the sequencing, bioinformatic analysis and functional validation of potential disease-causing variants identified within the trio WGS dataset of 100 Australian families impacted by neuromuscular disease who remain without a genetic diagnosis following best practice clinical genetic testing. The grant is also funding the development and testing of treatments targeting recessive titinopathy.

Dr Oates' UNSW Medical Genomics Group also has strong ongoing research collaborations with the Sydney Children's Hospitals Network, the Garvan Institute, the Victor Chang Cardiac Research Institute, as well as Sydney University, University of Western Australia, Murdoch University and Monash University. Her international collaborators include the Dubowitz Neuromuscular Centre (Great Ormond Steet Hospital/ University College London UK), Newcastle University



Clinical features of congenital titinopathy. Figure 2: Oates et. al. Congenital Titinopathy: Comprehensive characterization and pathogenic insights. Annals of Neurology 2018 Jun;83(6):1105-1124



Associate Professor Martin Smith Director, Ramaciotti Centre for Genomics

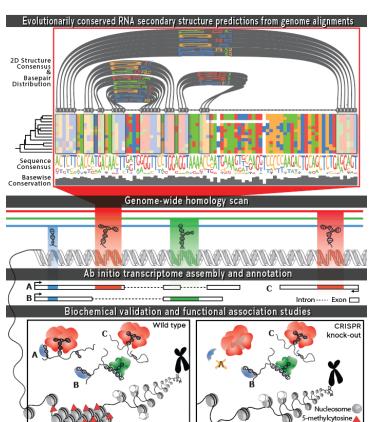
Room 2106, Level 2, Biological Sciences South Building (E26) E martin.smith@unsw.edu.au W therealsmithlab.com W unsw.edu.au/staff/martin-smith

RESEARCH PROGRAM

The Smith Laboratory develops cutting-edge bioinformatics tools to explore gene regulation using the latest genomic technologies. Positioned at the intersection of molecular cell biology and artificial intelligence, our research aims to transform health outcomes for complex diseases by unlocking the noncoding RNA code through innovative computational and experimental approaches.

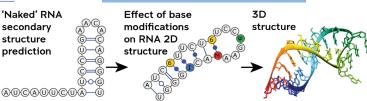
Our lab specializes in genomics and bioinformatics, with a strong focus on nanopore sequencing. We have led several pioneering developments in this space, including:

- the first high-throughput single-cell sequencing application of nanopore technology [1],
- the first end-to-end workflow for detecting RNA modifications [2],
- the first native RNA molecular barcoding method [3], and
- the first real-time RNA-based diagnostic workflow [4].



RESEARCH FOCUS

Computational Biology and Genomic Technologies



BIOGRAPHY

A/Prof Martin Smith directs the UNSW Ramaciotti Centre for Genomics, managing 26 professional staff, numerous robotics, genomics & computing instruments, and leads a research group of PhD students and postdocs in both Sydney and Montreal. He has over 19 years of genomics research experience across leading institutes in Canada, Germany, and Australia.

RESEARCH PROJECTS

We offer a range of research projects at the intersection of genomics, RNA biology, and artificial intelligence—each designed to push the boundaries of what's possible in modern biomedical science. Below are a few examples of current HDR opportunities, but we're always open to new ideas. If you're curious, motivated, and keen to explore how technology can unlock biology, feel free to get in touch to discuss tailored projects.

1. Molecular Dimensions of the Immune System

Enter the world of long-read and single-cell sequencing to chart the hidden dynamics of the human immune system. Work with patient data and nanopore sequencing to uncover how our defenses evolve in real time.

2. The Genome's Hidden Wiring

Unravel how a single genome builds complex life. Using Al and network theory, this project explores the deep regulatory circuits that guide mammalian development from the inside out via RNA structure.

3. The RNA Code: Nature's Logic Circuit

Is RNA the brain's secret code? Dive into the chemistry and computation of RNA logic, exploring neuromorphic computing and synthetic biology are powered by RNA structure and modifications.

4. Biomarkers That Matter

Hunt for molecular signals that define cancer and autoimmune diseases at unprecedented accuracy. Use advanced transcriptomics to build better diagnostics from the noise of the genome.

REFERENCES

- [1] Singh, M. et al. Nature Communications, 2019.
- [2] Liu, H. et al. Nature Communications, 2019.
- [3] Smith, M. A. et al. Genome Research, 2020.
- [4] Sagniez, M. et al. medRxiv, 2022.



Associate Professor Fatemeh Vafaee

Room 2110, Level 2 East Bioscience South Building E26 E f.vafaee@unsw.edu.au W https://vafaeelab.com/ W babs.unsw.edu.au/fatemeh-vafaee

RESEARCH FOCUS

Computational Biomedicine, Systems Biology, Bioinformatics.



CURRENT APPOINTMENT:

Deputy Director (Science), UNSW AI Institute (<u>UNSW.AI</u>), UNSW

Deputy Director, UNSW Data Science Hub (<u>uDASH</u>), UNSW Program Lead | Industry-Integrated Graduate Program (<u>Med-Tech.Al</u>)

Associate Professor, School of BABS, UNSW Sydney

BRIEF BIO AND RESEARCH CONTRIBUTION

Dr Vafaee is an Associate Professor at the School of Biotechnology and Biomolecular Sciences. She leads a multi-institute, industry-aligned next-generation graduate program, Med-Tech.Al on Al-enabled medical technologies, from diagnostics to therapeutics. She has established the Biomedical Al Laboratory (Vafaee Lab) at the UNSW Faculty of Science. The Biomedical Al Laboratory is a multidisciplinary team of researchers dedicated to developing sophisticated AI solutions in collaboration with industry and health organisations. Its work encompasses collaborative AI research to advance biomedical frontiers from disease diagnosis and drug discovery to single-cell multi-omics, with a commitment to precision, reproducibility, and interpretability. The team's efforts have led to significant translational achievements, moving technologies from proofof-concept to market readiness.

Recognised as one of the 'Top 10 Women in AI in APAC' by AI Magazine, Dr Vafaee also received the 2023 Women in AI Asia-Pacific Award in Health. She has also received the 2023 Mid-Career Research Contribution by Australian Bioinformatics And Computational Biology Society. Dr. Vafaee earned her PhD in Artificial Intelligence from the University of Illinois in Chicago and completed fellowships in computational biomedicine at the University of Toronto and the University of Sydney. With over a decade of experience, her R&D efforts have secured over millions in funding, leading to 60+ high-impact publications and the development of software tools, patents, and medical products currently on the path to commercialisation.

AREAS OF RESEARCH PROJECTS Theme 1: Al-Enhanced Disease Diagnostics and Personalized Medicine

New sequencing technologies enable the comprehensive analysis of various molecules (DNA, RNA, proteins) efficiently and affordably, generating vast amounts of omics data (genomics, transcriptomics, proteomics) from patient samples or individual cells. This data, alongside wearable sensor readings, electronic health records, and medical imaging, offers significant potential to advance personalized medicine and precision therapy. However, realizing this potential requires overcoming challenges through sophisticated AI models and big data analytics for effective data interpretation, prediction accuracy, and decision reliability. At the Biomedical Al Laboratory, we focus on improving disease diagnostics with generalizable models, enhancing non-invasive cancer management via better liquid biopsy tests, and increasing test sensitivity by incorporating multi-omics data and other patient information.

Theme 2: Single-Cell Multi-Omics: Tackling the Big Data Challenge

The advent of single-cell multi-modal omics (scMulti-omics) technologies has revolutionized our ability to measure diverse molecular features—like DNA methylation, chromatin accessibility, RNA expression, and protein levels-in individual cells, enabling a comprehensive understanding of cellular functions. Recognized by Nature Methods as the 2019 Method of the Year for its groundbreaking insights into cell functionality, novel cell-type discovery, and cross-omics relationships, the scMulti-omics field has rapidly grown. Despite this technological progress, the field faces significant computational challenges in integrating and interpreting the vast, complex datasets generated, hindering the accurate prediction of biological phenomena. Issues such as data heterogeneity, noise, systematic biases, and the curse of dimensionality are prevalent. At the Biomedical Al Laboratory, we employ AI to navigate these obstacles, enhancing the utility of single-cell data in life science research and clinical application.

Theme 3: Al-Enabled Drug Discovery

Al is transforming drug discovery and development, offering a revolutionary approach to the pharmaceutical industry. By analyzing complex datasets, including clinical, biological, molecular, and genetic information, Al algorithms enhance the precision and efficiency of drug development from drug target identification to toxicity assessments and compound design optimizations. Accordingly, Al facilitates the streamlining of research processes, from preclinical to clinical study designs, and supports the repurposing of drugs for new therapeutic applications.

We leverage advanced AI techniques to mine and interpret vast molecular, structural, and clinical datasets. This aids in repurposing drugs, identifying effective drug combinations, elucidating drug targets and mechanisms, and forecasting drug interactions and side effects, streamlining the path to novel therapeutics.

Selected Publications:

Besharatifard M; Vafaee F, 2024, 'A review on graph neural networks for predicting synergistic drug combinations', Artificial Intelligence Review, 57, http://dx.doi.org/10.1007/s10462-023-10669-z Zandavi SM et al, Vafaee F and Zekry A, 2024, 'Al-powered prediction of HCC recurrence after surgical resection: Personalised intervention opportunities using patient-specific risk factors', Liver International, http://dx.doi.org/10.1111/liv.16050 Vijayan A; Fatima S; Sowmya A; Vafaee F, 2022, 'Blood-based transcriptomic signature panel identification for cancer diagnosis: Benchmarking of feature extraction methods', Briefings in Bioinformatics, 23, http://dx.doi.org/10.1093/bib/bbac315 Zandavi SM; Koch FC; Vijayan A; Zanini F; Mora FV; Ortega DG; Vafaee F, 2022, 'Disentangling single-cell omics representation with a power spectral density-based feature extraction', Nucleic Acids Research, 50, pp. 5482 - 5492, http://dx.doi.org/10.1093/nar/ gkac436

Koch FC; Sutton GJ; Voineagu I; Vafaee F, 2021, 'Supervised application of internal validation measures to benchmark dimensionality reduction methods in scRNA-seq data', *Briefings in Bioinformatics*, 22, http://dx.doi.org/10.1093/bib/bbab304
Azad AKM; Dinarvand M; Nematollahi A; Swift J; Lutze-Mann L; Vafaee F, 2021, 'A comprehensive integrated drug similarity resource for in-silico drug repositioning and beyond', *Briefings in Bioinformatics*, 22, http://dx.doi.org/10.1093/bib/bbaa126





Professor Irina Voineagu Room 3107, Level 3 West Bioscience South Building E26 E i.voineagu@unsw.edu.au W babs.unsw.edu.au/irina-voineagu

RESEARCH FOCUS

Genetics of neurodevelopmental disorders, human brain transcriptome dynamics in normal and disease states.



RESEARCH PROGRAM

Broadly, my research interests are in the area of molecular genetic mechanisms underlying human brain disorders. My PhD work demonstrated that unstable DNA repeats block replication fork progression in bacteria, yeast and mammalian cells by forming DNA-hairpins on the lagging strand (Voineagu et al. PNAS 2008), and investigated for the first time the cellular checkpoint responses to replication fork arrest at CGG repeats. This work led to a novel model of chromosomal fragility at CGG repeat sequences (Voineagu et al. Nature Struct. Mol. Biol 2009). For postdoctoral research, I joined the Neurogenetics Department at UCLA, to investigate the molecular mechanisms of autism and intellectual disability, using transcriptome methods. My postdoctoral work led to the identification of a novel gene implicated in X-linked intellectual disability (Voineagu et al., Mol. Psychiatry 2011) and the characterisation of shared molecular pathways in autism post-mortem brain tissue (Voineagu et al. Nature 2011). Currently, my group's research concentrates on the molecular genetic mechanisms underlying normal brain function and their perturbation in neurodevelopmental disorders, using a combination of functional genomic studies in human brain tissue and neuronal cell culture systems.

More detailed information on projects and ongoing research is available on the lab website: www.voineagulab.unsw.edu.au

SELECTED PUBLICATIONS

- Won H, de la Torre-Ubieta L, Stein JL, Parikshak NN, Huang J, Opland CK, Gandal MJ, Sutton GJ, Hormozdiari F, Lu D, Lee C, Eskin E, Voineagu I, Ernst J, Geschwind DH (2016). Chromosome conformation elucidates regulatory relationships in developing human brain. *Nature* 538(7626):523-527
- » Yao P, Lin P, Gokoolparsadh A, Assareh A, Thang MW, Voineagu I (2015). Coexpression networks identify brain region-specific enhancer RNAs in the human brain. *Nature Neuroscience* 18(8):1168-74.
- » Voineagu I, Huang L, Winden K, Lazaro M, Haan E, Nelson J, McGaughran J, Nguyen L, Friend K, Hackett A, Field M, Gecz J, Geschwind DH (2012). CCDC22: a novel candidate gene for syndromic X-linked intellectual disability. *Molecular Psychiatry* 17(1):4-7.
- » Voineagu I, Wang X, Johnston P, Lowe JK, Tian Y, Horvath S, Mill J, Cantor RM, Blencowe BJ, Geschwind DH (2011). Transcriptomic analysis of autistic brain reveals convergent molecular pathology. *Nature* 474(7351):380-4.
- » Voineagu I, Surka CF, Shishkin AA, Krasilnikova MM, Mirkin SM (2009). Replisome stalling and stabilization at CGG repeats, which are responsible for chromosomal fragility. Nature Structural and Molecular Biology 16(2):226-8.
- » Voineagu I, Narayanan V, Lobachev KS, Mirkin SM (2008). Replication stalling at unstable inverted repeats: Interplay between DNA hairpins and fork stabilizing proteins. *Proc Natl Acad Sci U S A* 105(29):9936-41.



Associate Professor Paul Waters Room 3110, Level 3 East Bioscience South Building E26 E p.waters@unsw.edu.au W babs.unsw.edu.au/paul-waters



Sex chromosome structure, function, regulation and evolution.



My central area of expertise is sex chromosome biology and evolution, with a focus on epigenetic regulation of the X chromosome. In more recent years, I have also worked on hibernation in bearded dragons.

RESEARCH PROGRAM

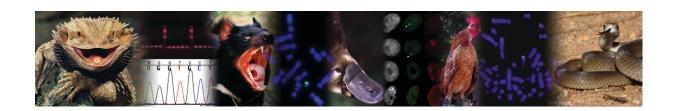
We use various molecular and bioinformatic methods to examining genome biology in diverse model species, unlocks answers about the evolution of complexities within our own genomes. The lab focuses on better understanding the epigenetic regulation of transcription in distantly related vertebrate representatives, specifically focusing on sex chromosomes. The ultimate goal is to understand how complex epigenetic silencing mechanisms evolved. We use eutherian, marsupial, monotreme and bird/reptile models. Representatives from these groups each have different (sometimes weird and wonderful) sex determining mechanisms and sex chromosome systems.

We have a keen interest in X chromosome inactivation and meiotic sex chromosome inactivation in marsupial models. We also have particular interest in dosage compensation of the strange sex chromosomes of platypus, which have 5 X chromosomes and 5 Y chromosomes! We also study the Australian central bearded dragon, which are unusual in that they have both genetic and environmental sex determination. We also have a lot of interest in 3D genome structure, especially in the transmissible Tasmanian devil facial tumour.

Click here for a full list of publications.

SUPERVISION

PhD, MSc, Honours, and 3rd year Undergraduate projects in Epigenetics and Molecular Biology are available in the lab.







Associate Professor Robert Weatheritt

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RESEARCH FOCUS

RNA and phenotypic complexity: how regulation of RNA impact protein function; alternative splicing; membraneless organelles; CRISPR single cell screens.



RESEARCH PROGRAM

An important question in biology is how the complexity of biological systems has expanded while the number of protein coding genes has remained largely stable. Decades of research has shown that increased biological complexity has arisen in part by the dynamic generation of unique, cell-specific transcriptomes, as well as the dynamic subcellular control of these transcripts. Alternative splicing (AS) - the process by which multiple, distinct transcript and protein variants are expressed from a single gene - is a key driver of transcriptome complexity regulating more than 95% of multi-exon genes. However, the functional flexibility and complexity afforded by AS, and other post-transcriptional regulation, comes at a cost, since it increases the probability that RNA mis-splicing may arise and lead to human disease. By developing novel state-of-the-art genomic tools and large-scale screening approaches, my work has revealed instances in which splicing misregulation impacts both neurodevelopmental and somatic disorders.

RESEARCH PROJECTS

Project 1: Alternative splicing and membraneless organelles in neuronal differentiation. This project will investigate the role of RNA in regulating the function of membraneless organelles and its role in controlling neuronal differentiation, as well as its implications on neurodevelopmental disorders.

Project 2: Membraneless organelles and chemoresistance. This project will aim to understand the role of stress granules and P-bodies in chemoresistance.

Project 3: Developing deep learning algorithms to discern impact of epitranscriptomic modifications on the genome. This project will develop algorithms to understand the genetic code controlling RNA modifications identified by nanopore long-read data.

BIOGRAPHY

Associate Professor Weatheritt is an EMBL Australia group leader and an ARC Future Fellow. Robert received his PhD from the European Molecular Biology Laboratory in Heidelberg, Germany. He completed his postdoctoral training in Cambridge and Toronto before beginning his lab at the Garvan Institute in 2018. Since 2022, Robert has a joint appointment at School of Biotechnology and Biomolecular Sciences.

Contact Robert about research supervision opportunities at: r.weatheritt@unsw.edu.au

RELEVANT RECENT PUBLICATIONS

Evolution of splicing:

- "Evolutionary dynamics of circular rnas in primates" ELife (2021)
- "Multilayered control of exon acquisition permits the emergence of novel forms of regulatory control" Genome Biology (2019)

Membraneless organelles:

- "Systematic mapping of nuclear domain-associated transcripts reveals speckles and lamina as hubs of functionally distinct retained introns" Molecular Cell (2022)
- "Regulatory expansion in mammals of multivalent hnRNP assemblies that globally control alternative splicing" Cell (2017)

Splicing and neuroscience:

- "Differential contribution of transcriptomic regulatory layers in the definition of neuronal identity" Nature Communications (2021)
- "Autism-misregulated eIF4G microexons control synaptic translation and higher order cognitive functions" Molecular Cell (2020)
- "A highly conserved program of neuronal microexons is misregulated in autistic brains". Cell (2014)

Splicing algorithms:

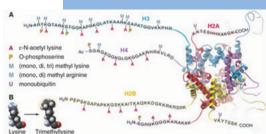
- "Efficient and accurate quantitative profiling of alternative splicing patterns of any complexity on a laptop" Molecular Cell (2018)
- "The ribosome-engaged landscape of alternative splicing" Nature SMB (2016)



Professor Marc Wilkins Deputy Dean (Research & Enterprise) Room 2112, Level 2 East Bioscience South Building E26 E marc.wilkins@unsw.edu.au W babs.unsw.edu.au/marc-wilkins

RESEARCH FOCUS

Discovery and functional characterisation of intracellular networks.



RESEARCH

My research questions are centred around proteomics.

1. What is the regulatory network of histone methylation? Histone methylation is a crucial process that affects the compaction and relaxation of chromatin, and thus gene expression in the cell. The sites of histone methylation are understood and the enzymes responsible are known, at least in the model system of yeast. However we know little about how the four 'writer' enzymes and the four 'eraser' enzymes are actually regulated and how they work as a

- Are the histone methyltransferases and demethylases phosphorylated?
- If phosphorylation on these enzymes affect their activity and if so, how?
- What kinases are responsible for this phosphorylation?

single integrated system. We are exploring:

 Do the modification of writer and eraser enzyme control where they act, on chromatin in the genome?

We aim to connect the cell's signalling system with its histone-based system of gene regulation in this project.

The project is supported by ARC Discovery Project grant 2020: The Regulatory Network of Histone Methylating and Demethylating Enzymes.

Recent papers on protein methylation:

- Characterization of Protein Methyltransferases Rkm1, Rkm4, Efm4, Efm7, Set5 and Hmt1 Reveals Extensive Post-Translational Modification. Winter DL, Hart-Smith G, Wilkins MR. J Mol Biol. 2018 430(1):102-118.
- Methylation of Elongation Factor 1A: Where, Who, and Why? Hamey JJ, Wilkins MR. Trends Biochem Sci. 2018 43(3):211-223.
- METTL21B Is a Novel Human Lysine Methyltransferase of Translation Elongation Factor 1A: Discovery by CRISPR/Cas9 Knockout. Hamey JJ, Wienert B, Quinlan KGR, Wilkins MR. Mol Cell Proteomics. 2017 Dec:16(12):2229-2242.

2. How do cells make decisions via protein interaction networks?

Cells use protein post-translational modifications as an 'information management' and 'status management' system. Yet how this information is integrated, to make actual decisions inside networks, is poorly understood.

We are exploring:

- How often are two post-translational modifications found next to each other in the proteome? Especially methylation, phosphorylation and acetylation?
- For modifications that are nearby, what is their 'logic'
 do they block each other? Is one required before the other?

- Do nearby modifications form small 'decision making modules' in interaction networks?
- Do nearby modifications change the interaction choice of a protein? Or change its localisation in the cell?

The project is supported by ARC Discovery Project grant 2017: The discovery of decision-making modules in protein interaction networks.

Recent papers on crosstalk of post-translational modifications:

- Crosstalk of Phosphorylation and Arginine Methylation in Disordered SRGG Repeats of Saccharomyces cerevisiae Fibrillarin and Its Association with Nucleolar Localization. Smith DL, Erce MA, Lai YW, Tomasetig F, Hart-Smith G, Hamey JJ, Wilkins MR. J Mol Biol. 2020 432(2):448-466.
- Knockout of the Hmt1p Arginine Methyltransferase in Saccharomyces cerevisiae Leads to the Dysregulation of Phosphate-associated Genes and Processes. Chia SZ, Lai YW, Yagoub D, Lev S, Hamey JJ, Pang CNI, Desmarini D, Chen Z, Djordjevic JT, Erce MA, Hart-Smith G, Wilkins MR. Mol Cell Proteomics. 2018 17(12): 2462-2479.

3. How is the protein interactome regulated?

Recent, breakthroughs in crosslinking mass spectrometry (XL-MS) have made it possible to measure thousands of protein-protein interactions in a single sample. These generate a 'protein interactome' which reflects the biological state of a system, at a point in time. Use of heavy isotope-based techniques with XL-MS allows 'protein interactomes' to then be compared. We are exploring:

- The optimisation of techniques for XL-MS.
- The roles of gene expression and protein posttranslational modifications in the regulation of the protein interactome.
- The role of alternate splicing of mRNA and resulting isoforms in regulating the protein interactome.

Recent papers on crosslinking mass spectrometry:

- Cross-linking Mass Spectrometry Analysis of the Yeast Nucleus Reveals Extensive Protein-Protein Interactions Not Detected by Systematic Two-Hybrid or Affinity Purification-Mass Spectrometry. Bartolec TK, Smith DL, Pang CNI, Xu YD, Hamey JJ, Wilkins MR. Anal Chem. 2020 92(2):1874-1882.
- Characterization of the Interaction between Arginine Methyltransferase Hmt1 and Its Substrate Npl3: Use of Multiple Cross-Linkers, Mass Spectrometric Approaches, and Software Platforms. Smith DL, Götze M, Bartolec TK, Hart-Smith G, Wilkins MR. *Anal Chem.* 2018 90(15):9101-9108.

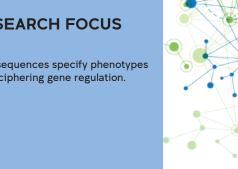




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RESEARCH FOCUS

How sequences specify phenotypes by deciphering gene regulation.



The human genome contains roughly 20,000 proteincoding genes but hundreds of thousands of regions responsible for tuning the activation of these genes in space and time. We are interested in the interplay between these regions and the circuits they control. Those interactions between the genome and the epigenome ultimately specify cell diversity and animal form and function. Our lab uses computational and statistical methods, and evolutionary concepts to generate hypotheses and interrogate data. We are largely computational, but we also go beyond the dry lab to generate molecular data to address fundamental biological and biomedical questions.

BIOGRAPHY

A/Prof Emily Wong received her Masters and PhD from the University of Sydney in Bioinformatics and Computational Genomics. She was an EMBO postdoctoral fellow at the European Molecular Biology Laboratory-European Bioinformatics Institute (EMBL-EBI) at the Wellcome Trust Genome Campus, Cambridge, UK. There, she studied the regulatory evolution of mammalian tissues. In 2016, She returned to Australia as an Australian Research Council Discovery Early Career Fellow to the University of Queensland where she worked on genomic control during development.

Emily is part of UNSW and the Victor Chang Cardiac Research Institute. Her research seeks to tie together genetic and molecular understanding to decipher the rules controlling cell and trait diversity. She uses new cross-disciplinary approaches integrating big data with in vivo experiments. Her work leverages the power of comparative genomics to understand how traits are encoded in our genomes and how regulatory systems are disrupted in disease.

RECENT PUBLICATIONS

- Wong ES*,...Francois F*, Degnan B* (2020). Deep conservation of the enhancer regulatory code in animals. Science *corresponding
- Flochay S^, ES Wong^, Zhao B^... Garfield D, Furlong E (2021). Cis-acting variation is common, can propagate across multiple regulatory layers, but is often buffered in developmental programs Genome Research ^ equal contributions
- Wong ES, Schmitt B, Thybert D, Marioni J, Ferguson-Smith A, Odom DS, Flicek P (2017) Interplay of cis and trans mechanisms driving transcription factor binding, chromatin, and gene expression evolution. Nature Communications 8(1):1092
- Wong ES, Thybert D, Schmitt B, Stefflova K, Odom DS, Flicek P (2015) Decoupling of evolutionary changes in transcription factor binding and gene expression in mammals. Genome Research 25:167-78

RESEARCH SUPERVISORS -

MICROBIOLOGY AND MICROBIOMES

CLUSTER STRENGTHS:

- » Microbes in Health and Disease
- » Microbes in the Environment

Microbes are invisible companions that intertwine our biology and support our biological and geological systems. They are big players in infectious diseases but are also fundamental to producing nutrients for plants to grow and the dynamic transformation of matter. We aim to unravel the mechanisms behind these ubiquitous microbes and their vital function in every life process. Our research in Microbiology & Microbiomes explores the importance of microbes in the environment and microbial contributions to health and disease.

Our students are encouraged to use their critical and analytical aptitude and exercise a range of genomic tools to address global topics such as archaea, climate change and food production. We endeavour to translate our research into effective methods for the control and treatment of conditions like autism, cancer and diabetes. Driven by improvements in technology and the imaginations of our researchers, we aspire to unravel the many secrets of the microbial world.



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Environmental microbiology (microbial diversity, adaptation, evolution, ecosystem function) and astrobiology (early life and human health)

Our research is focused on unravelling the evolutionary and ecological significance of early Earth microbial ecosystems.

Stromatolites and microbial mats are model systems for studying the origins and evolution of life on our planet. They are geobiological structures composed of complex and diverse microbial communities. We have access to unique field sites on the coast of Western Australia – in particular the World Heritage site of Shark Bay - and other locations around the world. We also work closely with the Department of Parks and Wildlife to ensure these unique ecosystems are carefully monitored in the face of threats such as climate change. In particular, the impact of extreme stressors on microbial communities and critical pathways in threatened mat systems are being assessed and critical to ascertain before any irreversible ecosystem tipping points are reached.

The study of microorganisms associated with these formations may also be applied to the search of extraterrestrial life (past or extinct), particularly with the discovery of unique bio-signatures. This work thus aligns well with the goals of the Australian Centre of Astrobiology and our collaborators at NASA. Our research provides new metagenome-based models into how biogeochemical cycles and adaptive responses may be partitioned in the microbial mats of Shark Bay, including the genetic basis for novel natural product synthesis. The traditional tree of life is also in flux, and new discoveries we are making of novel organisms and pathways is affording a dynamic and holistic view of these ecosystems.

In particular, we are pursuing the role of 'microbial dark matter' in these systems including the enigmatic group of Asgard archaea. We aim to break down the traditional distinctions between prokaryotic and eukaryotic life using the Asgardians as a 'missing link'.

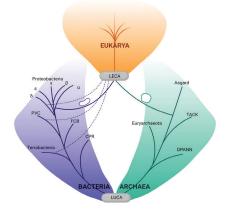
This research combines biogeochemical field measurements, laboratory analytical methods, and recent advances in functional genomics. In particular, there is the opportunity to employ next-generation sequencing platforms, including various 'meta' approaches (genomics, transcriptomics, proteomics). Students will use these and other modern microbial and molecular biology techniques to examine specific aspects of community function in these 'living rocks', from deciphering microbial interactive networks, novel adaptive responses and natural product synthesis.

Specific project areas include:

- Exploring the unknown: illuminating microbial dark matter in mats
- Promiscuity in microbial mat communities: gene transfer and impact of viruses
- Searching for our great (cellular) ancestors: hunting the elusive Asgard archaea
- The canary in the coalmine: effects of environmental change on microbial communities
- Living at the edge: understanding microbial survival in an extreme environment
- Look who's talking too: communication in the third domain of life
- Mining for novel natural products: microbial mats as a source for unique metabolites

I also encourage students who want to think outside the box, so I always welcome ideas for other projects and happy to workshop potential!







Associate Professor Natalia Castaño-Rodríguez NHMRC Emerging Leadership Fellow (EL2) and UNSW Scientia A/Prof

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RESEARCH FOCUS

Gastrointestinal disease.

BIOGRAPHY AND RESEARCH CONTRIBUTION

A/Prof Natalia Castaño Rodríguez is a WSTEM Early Career Researcher, who has worked in the fields of molecular biology, microbiology, biostatistics and bioinformatics related to immunogenetics and genetics of cancer for the past 12 years. She currently leads a research group composed of higher degree research and Honours students, all dedicated to investigate the interplay between host immunogenetic factors and gastrointestinal dysbiosis in gastrointestinal carcinogenesis (mainly stomach cancer) and other chronic inflammatory conditions (mainly inflammatory bowel diseases).

Prior to moving to Australia, Natalia undertook research trainee fellowships in Colombia (Biological Research Corporation (CIB), Medellín, Colombia) and Mexico (Occidental Biomedical Research Centre (CIBO), Guadalajara, Mexico), where she acquired skills in the areas of human genetics and autoimmunity.

Following the completion of her MD degree, Natalia funded her postgraduate studies with competitive scholarships from Colombia and Australia (COLFUTURO Scholarship and UNSW University International Postgraduate Award) to investigate the role of innate immunity in stomach cancer. After a stint as a research associate in the Helicobacter and Campylobacter laboratory at UNSW, she was then awarded an Early Career Fellowship by the National Health and Medical Research Council (NHMRC) to expand her work on the role of immunogenetics, particularly related to pattern recognition receptors and the autophagy pathway, in Helicobacter pylori-related stomach cancer (May 2016- April 2020). During this time, she further funded her research with a Cancer Australia Priority-driven Collaborative Cancer Research Scheme (PdCCRS) Young Investigator grant (2017) as well as a Gastroenterological Society of Australia Project Grant (2019) to investigate the contribution of dysregulated autophagy to carcinogenesis in the stomach. Having established herself as an emerging leader in the field of immunogenetics in cancer, Natalia recently secured a Cancer Institute NSW ECF (2020-2023) as well as a UNSW Scientia Fellowship (2020-2024).

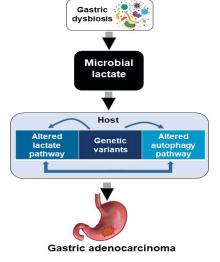
If you would like to learn more about Natalia and her research, please read these layperson articles published in <u>Nature Communications Biology</u>, <u>Can Too Foundation</u> and <u>Medscape</u>. A full list of Natalia's scientific publications can also be found here.



Figure 1. Gastric organoids. Example of an organoid growing from a human gastric biopsy in our laboratory at days 0, 4, 7 and 9.

RESEARCH PROGRAM

Stomach cancer is the 5th most common cancer and the 3rd leading cause of cancer-related mortality worldwide, with incidence rates particularly high in East Asia and South America. Notably, stomach cancer incidence is equally high in Indigenous Australians while the incidence in non-Indigenous Australians will increase due to an



ageing population and high influx of migrants from East Asia and South America, placing stomach cancer as a research priority in our nation. While Helicobacter pylori is a fundamental risk factor, stomach cancer aetiology involves the combined effects of microbial, host and environmental factors. This is evidenced by the fact that only 1-3% of H. pylori infected people develop stomach cancer, and that progression to stomach cancer in some subjects occurs even after eradication of the bacterium. Natalia proposes a synergistic interaction between innate immunity (mainly pattern recognition receptors (PRRs) and autophagy), H. pylori infection and gastric dysbiosis in stomach cancer. Based on this hypothesis, her research program can be divided into the following arms: 1) Autophagy in gastric carcinogenesis, 2) The role of the gastric microbiome (H. pylori and non-H. pylori taxa) in gastric carcinogenesis, 3) The role of microbial metabolites in gastric carcinogenesis, and 4) Pattern recognition receptors in gastric carcinogenesis.

AREAS OF SUPERVISION

Natalia is interested in supervising both Honours and HDR (Masters and PhD) students in the areas of immunology, genetics and microbiology. If you are interested, please contact her via email describing why you would like to undertake a degree in her lab.



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Exploring microbial dark matter and microbial processes in cold desert environments.

RESEARCH GOALS

- Determine the resilience of Antarctic soil microbial communities to environmental change
- To uncover the global significance of atmospheric chemosynthesis and identify the potential for microbial life elsewhere
- To develop novel cultivation approaches for yet-tobe cultured microbes and fill gaps in the tree of life
- Develop ecotoxicity assays for hydrocarbon contamination in cold regions using microbes as indicators of soil health
- Explore novel natural products in extreme bacteria and fungi

RESEARCH IN DETAIL

I have built up strong partnerships across both the Biotechnology industry and government bodies in Australia. My research has real-world applications, driving remediation targets, guideline derivation and conservation efforts in Antarctica.

In Antarctic soils, microbes are the most dominant lifeform and thus they drive geochemical processes, particularly carbon and nitrogen cycling. My research is aimed at unravelling the breadth of microbial diversity and their functioning in soil. My team focuses on microbial dark matter, that is bacteria, archaea and fungi that are yet-to-be cultured and we have recently isolated a suite of novel fungi that now require characterisation.

By integrating with genomics with proteomics and multivariate analyses, my group is exploring the ecology of microbes in both pristine and contaminated soils. Through collaboration with the Australian Antarctic Division, the Botanic Gardens and NASA we are using molecular tools to determine the limits for life, and to evaluate soil health in response to natural and man-made disturbances, from hydrocarbon contamination through to climate induced change. My research is world-class, and of high impact, with our discovery of Antarctica bacteria surviving by literally 'living on thin air' published in the journal Nature. My research is challenging our understanding of the nutritional limits required to support life and has opened up the possibility for life on other planets.

Please see <u>Ferrarilab.org</u> for more details including information on our 2019 expedition to the Windmill Islands, east Antarctica.

ABOUT THE LAB HEAD

I am a Professor in Environmental Microbiology and am currently the Associate Dean Research in the Faculty of Science. I specialise in Antarctic soil biodiversity and perform discovery-based and applied research. I am passionate about training the next generation of confident scientists, being awarded the Vice-Chancellors Award for Excellence in Higher Degree Research Supervision in 2021. I have supervised >20 Honours and 23 HDRs to completion with my student's gaining employment across both academia and industry. Through my collaborations with the Australian Antarctic Division, the Botanic Gardens and NASA there are opportunities for Industry-associated PhDs in my team.

ABOUT THE LAB

My research team is currently comprised of three postdoctoral scientists, one part time research associate, six PhD and two Honours students. We have a supportive environment that is well-known for providing strong mentorship to students throughout their candidature. We have a range of wet- and dry lab projects, spanning the characterisation of novel bacteria and fungi, the isolation of new natural products, through to the development of ecotoxicity assays and assessments on contaminated soils in Antarctica, and finally exploring the significance of atmospheric chemosynthesis to primary production in desert soils.



Figure 1. Mitchell Peninsula, Antarctica; a nutrient-limited desert that hosts a unique microbial community that uses trace gases to survive.



Figure 2. Casey station where bioremediation of fuel spills is ongoing using engineered biopiles combined with nutrient amendment.



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Genomics and molecular evolution of bacterial pathogens.



Infectious diseases caused by pathogenic bacteria are a major threat to human health. Our group takes a multi-disciplinary approach to study pathogenic bacteria. We work on a number of bacterial pathogens including Bordetella pertussis, Salmonella, Shigella and Vibrio cholerae. We use omics (genomics, transcriptomics, and proteomics) approaches to address how pathogens arise and cause disease, how they evolve and adapt and how to identify these pathogens. These studies are significant in designing strategies that will be effective in preventing the emergence and spread of pathogens.

RESEARCH PROGRAM

Evolution and virulence of respiratory tract pathogen *B. pertussis*

Pertussis, commonly known as whooping cough, is an acute respiratory disease caused by B. pertussis. Despite widespread vaccination, pertussis remains a public health burden. Australia is currently experiencing a prolonged pertussis epidemic, with nearly 40,000 cases at its peak in 2011 and the second from 2014 to 2017 with 20,000 cases at its peak. Recent pertussis epidemics have also been observed in other countries with high vaccine coverage and several causes have been suggested for the resurgence of pertussis. Our research has provided strong evidence that adaptation of *B. pertussis* in response to selection pressure from pertussis vaccines has contributed to the pertussis resurgence. In this program we aim to understand how B. pertussis involve under the selection pressure of vaccines and antibiotics. We also aim to understand the virulence and pathobiology of *B. pertussis* using both genomic and proteomic approaches to improve the current vaccine.

Genomic epidemiology, surveillance, and evolution of bacterial pathogens

The effectiveness of public health interventions for infectious disease control is limited by the low resolution of current surveillance methods and a limited understanding of pathogen evolution. Both can be radically improved by whole genome sequencing (WGS), which enables the identification of outbreak cases, detection and tracking of emerging epidemic clones and elucidation of the factors that contributed to their emergence and spread. This research program aims to develop novel methods and solutions to enhance infection control and prevention through genomics guided surveillance.

We pioneered a new approach for genomic surveillance of Salmonella Typhimurium. We have developed multi-level genome typing (MGT), using multiple, sequential multilocus sequence typing (MLST) schemes of increasing size that allow examination of genetic relatedness at resolutions from seven gene MLST to core genome (cg) MLST. MGT provides the best means to identify strains and genome types at a resolution appropriate to the needs of short and long term epidemiology. We implemented this system for STM to demonstrate its utility and an online MGT database (http://mgtdb.unsw.edu.au) for public health applications. We aim to apply similar approaches to other bacterial pathogens for outbreak detection and epidemiological surveillance for better control of outbreaks, disease spread and antibiotic resistance. We also aim to understand pathogen evolution and adaptation through novel bioinformatic and genomic analyses.

ABOUT THE LAB HEAD

I am a Professor of Medical Microbiology in the School of Biotechnology and Biomolecular Sciences at UNSW Sydney where I teach medical microbiology to science and medical students. I grew up in the southeast countryside of China and did my undergraduate degree in China. I moved to Australia in 1986. I completed my PhD at University of Sydney in 1992 and did my postdoctoral training also at the University of Sydney. I was appointed as a Senior Lecturer at UNSW in 2002 and promoted to Professor in 2018. I have published over 190 papers with >10,000 citations (Google scholar). I invented the core genome concept which is central to the understanding of bacterial evolution.

ABOUT THE LAB

Our research group currently has 1 postdoctoral associate, 1 Bioinformatician, 6 PhD students and 2 honours students. Our Lab is known for its collaborative and supportive research environment that encourages innovation and creativity to unlock each student's potential. We welcome students from around the world with academic backgrounds in microbiology, genetics, bioinformatics, and related disciplines.

Our past and present students came from around the world and our graduates have gone on to successful careers including university professors, senior scientists, and laboratory heads.

For more information about us please visit our lab website http://www.lanlab.unsw.edu.au.



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Fungal infections of humans.

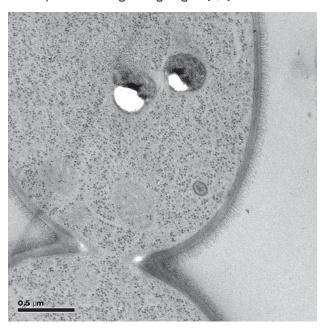
Opportunistic invasive fungal pathogens cause over 6.5 million life-threatening infections and 3.75 million deaths per year worldwide. Bloodstream and invasive infections caused by *Candida* species are one of the most frequent life-threatening invasive fungal infections. Almost half of the 1.5 million cases of invasive candidiasis per year are caused by one species, *Candida albicans*, which is one of the World Health Organisation's critical group fungal priority pathogens.

C. albicans colonises the gastrointestinal (GI) tract of around 80% of healthy individuals. It persists as a harmless commensal when kept in check by competition with the bacterial microbiota, innate and adaptive immune functions and gut epithelial cells. Invasive infections occur when these mechanisms are impaired (e.g. in the seriously ill and immunocompromised). Even when diagnosed and treated, more than 35% of people who get these infections die.

Research in my group is aimed at finding new ways to treat or prevent invasive *C. albicans* infections.

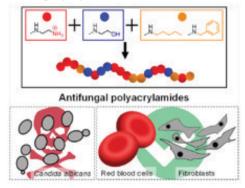
Fungal cell wall structure and biosynthesis:

We study fungal cell wall structure and biosynthesis to identify new antifungal drug targets [1,2].



Novel antifungal polymers:

We are working with chemical engineers to develop novel antifungal polymers [3,4].

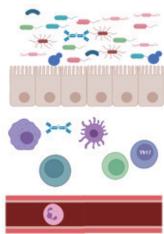


Microbiome-based interventions:

We also study how *C. albicans* colonises the GI tract with the aim of developing microbiome-based interventions to prevent invasive *C. albicans* infections [5,6].

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- 5. Ricci et al. (2022). Human gut bifidobacteria inhibit the growth of the opportunistic fungal pathogen *Candida albicans. FEMS Microbiology Ecology* **98(10)**:fiac095.
- 6. McCrory, Lenardon and Traven (2024). Bacteriaderived short-chain fatty acids as potential regulators of fungal commensalism and pathogenesis. *Trends in Microbiology:* **32**:1106-1118.





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RESEARCH FOCUS

Essential bacterial colonisation factors

My research group aims to understand how bacterial pathogens cause disease in order to develop new treatments and/or vaccines. We are particularly interested in understanding what are the essential colonisation and virulence factors that different bacterial pathogens need to infect hosts. By targeting essential colonisation factors, we can develop new treatments and vaccines which limit the bacteria's ability to develop resistance or escape.

My group focuses on two pathogens: Bordetella pertussis (Whooping cough) and Chlamydia trachomatis. We use a combination of omics (genomics, transcriptomics, proteomics) and cuttingedge molecular biology techniques (TraDIS, CRISPRi, cloning, qPCR etc.) to dissect host-pathogen interactions.

Project 1: T cell antigen discovery for Bordetella pertussis:

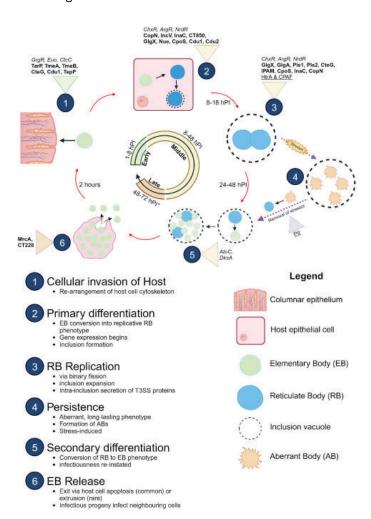
Bordetella pertussis causes whooping cough, one of the most prevalent vaccine-preventable disease with >23,000 cases in 2024. One reason for the remergence of whooping cough is the inability for the current vaccine to stimulate a Th1/Th17 response, which has been shown to be essential for protection. To correct this, this project aims to discover new and validate pertussis T-cell antigens using immunoproteomics to improve the current vaccine.

Project 2: Essential genes for biofilm and survival of Bordetella pertussis cells in microaerobic conditions

B. pertussis is widely thought to be an obligate aerobic pathogen. However, we have shown that not only can *B. pertussis* survive in microaerobic environments, it actually forms increased biofilms under low oxygen conditions such as those found in the lungs. We want to investigate this phenomenon further and uncover the mechanism behind *B. pertussis* ability to thrive in microaerobic environments.

Project 3: Understanding the asynchronous lifecycle of Chlamydia

Chlamydia trachomatis is the most common sexually transmitted bacteria with over >100,000 cases each year. It is a strict intracellular bacteria that only reproduces inside human cells. During infection, this pathogen asynchronously switches from reticulate bodies (RB), which can divide but cannot infect, to elementary bodies (EB), which can infect but cannot replicate. We are interested in understanding the factors governing the asynchronous conversion of RB to EBs. Understanding how Chlamydia progresses through its development cycle will allow us to develop new treatments targeting RB to EB conversion, thereby blocking infection.





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Evolution of pathogens: computational biology and mathematical modelling.

RESEARCH AREAS

In our group we are interested in understanding how microbes evolve. We use mathematical, statistical and computational methods to model the population dynamics of pathogens and other microorganisms and to analyse genetic data. Recent research areas include the dynamics of gut microbial organisms, microbial niche construction, the evolution of pathogens across multiple scales, the role of cultural evolution in pathogen evolution, and the extinction of asexual populations. The following are some current questions.

How old is a pathogen? Genomes contain information about the history of a species. For example, when free-living bacteria turn into pathogens they undergo reductive evolution and relaxed selection on some genes which then become pseudogenes. One way to infer the age of a pathogen is by studying patterns of molecular evolution in genes and pseudogenes; this was done to estimate the age of leprosy, for instance. This project aims to refine the methodology for this estimation procedure and to apply it to a variety of bacterial pathogens.

How do technology and culture influence the transmission of infectious diseases? Technological advances have opened many opportunities for economic development but they have simultaneously altered the way we interact with the environment. For instance, we have increased contact with wildlife and potentially raised the risk of zoonoses. This project aims to study how the dynamics of human cultural practices contribute to this risk.

How do pathogens adapt or go extinct? When pathogens spread in host populations during epidemics, their population sizes expand and contract. They may go extinct after the contraction or cause further outbreaks. Such population dynamics can be captured by mathematical models. Although these models have been well studied it is less clear how evolution occurs in oscillating populations. While population genetic models have been well-studied, they focus on constant population sizes. This project aims to predict the trajectories and consequences of new mutations in oscillating pathogen populations.

For more information about us please visit our lab website: www.tanakalab.unsw.edu.au

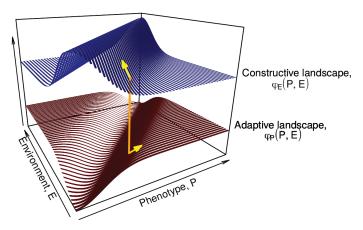


Figure: The Dual Landscape model of adaptation and niche construction. Adaptive landscape (lower surface) and constructive landscape (upper surface). Vectors show partial change in the directions of the phenotype ${\cal P}$ on the lower landscape and the environment ${\cal E}$ on the upper landscape. The pole connecting the two vectors shows that they are located at corresponding positions in the ${\cal P}\text{-}{\cal E}$ space.



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Gene regulation in bacterial pathogens.

The group is interested in the molecular basis of pathogenesis in *E. coli* and multidrug resistant *Staphylococcus aureus* (MRSA). These pathogens are significant causes of human morbidity and mortality and understanding how they cause disease is an important step towards developing new therapeutics.

The lab is particularly interested in how these pathogens control virulence and antibiotic resistance genes. We have made important contributions to understanding the functions of non-coding RNAs (ncRNAs) in these pathogens and have identified ncRNAs that control antibiotic tolerance and toxin expression.

RESEARCH PROGRAM

Antibiotic tolerance in *Staphylococcus aureus* (MRSA) Staphylococcus aureus is one of the most adaptable and frequent human pathogens, causing disease at almost every site in the body. Antibiotic resistance is a major problem and mortality rates for patients with MRSA septicaemia are 15-50%. Last-line antibiotics are used to treat these infections, but antibiotic tolerance is increasingly causing treatment failure.

We work to understand the molecular basis of antibiotic tolerance in MRSA and have recently identified previously unknown genes and ncRNAs that contribute to tolerance.

Virulence in toxigenic *E. coli*

Shiga-toxigenic *E. coli* cause potentially fatal kidney and neurological disease. Antibiotic treatment of these infections is not recommended as treatment can increase toxin release. We have identified many new ncRNAs that control virulence and toxin expression in toxigenic *E. coli*, potentially providing new avenues for treatment.

Enhancing antisense RNA therapeutics

We are applying our understanding of ncRNA function in pathogenic bacteria to enhance the activity of antisense RNA therapeutics. These molecules can be used as sequence-specific antibiotics or anti-virulence compounds and may represent a promising platform technology to combat antimicrobial resistance.

ABOUT THE LAB

We are a diverse and supportive lab committed to providing students with a platform to achieve their best. We have an excellent balance of senior researchers (post-docs), PhDs, and Honours students which provide an outstanding support network and make the lab energetic and fun. More information can be found at treelab.science.

ABOUT THE LAB HEAD

I am an Associate Professor of Microbiology. I obtained my PhD in Microbiology at the University of Queensland and was a post-doctoral researcher at the University of Edinburgh (UK) and the University of Melbourne before moving to UNSW in 2015. I enjoy finding the biological meaning within complex datasets and understanding how life (and bacteria!) work at a molecular level.





Bacteriophage are common vectors for transferring virulence genes between bacteria.



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RESEARCH AREAS

Viral infections are a major global health threat and burden. Unless we understand the mechanisms, patterns, and consequences of their rapid evolution, we will not be able to build rational strategies for controlling their spread. Our research encompasses molecular virology, viral discovery, molecular surveillance, drug discovery, host-virus evolution and tickborne disease to better understand what makes a successful virus. We use a multi-disciplinary approach in our projects, combining wet-lab molecular biology and microbiology techniques with computational analysis and bioinformatics.

SPECIFIC PROJECTS

Our research revolves around four main areas.

Molecular epidemiology of human enteric viruses

All human pathogens regularly circulate within the community, genetically evolve and emerge to cause outbreaks. Norovirus, a leading cause of acute gastroenteritis worldwide, it regularly attacks cruise ship, aged-care centres, hospitals, and childcare. Pandemic strains can take this to a global scale with hundreds of millions of cases of acute gastroenteritis yearly when a pandemic strain strikes, as in 2013, caused by the Sydney 2012 norovirus which was discovered by our group at UNSW.

In our research we use a combination of clinical and wastewater samples to monitor the evolution of norovirus strains over time, identify recombination events and antigenic shift, and link these molecular changes to epidemiological data. We also conduct similar monitoring on human adenovirus and SARS-CoV-2.

Viral discovery

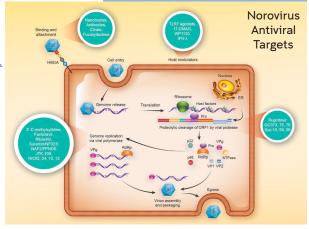
Over 70% of pandemics are caused by zoonotic viruses; HIV, SARS-CoV-2, and Zika virus are some examples from recent years. Using a combination of RNA-sequencing and bioinformatics, we discover viruses in a range of animals including bats and ticks. We also aim to discover viruses in vulnerable and endangered populations to aid in the detection and management of future viral outbreaks.

Endogenous viral elements in host genomes (paleovirology)

Endogenous viral elements (EVEs) are viral "fossils" inside host DNA. These fossils can be used to study ancient viruses and understand virus evolution over millions of years. We study the prevalence and diversity of EVEs throughout vertebrates and invertebrates to better understand the age and genetic evolution of

RESEARCH FOCUS

Molecular virology.



viral families. In addition, we have evidence that EVEs contribute to cellular function and therefore they can protect cells from infection.

Antiviral discovery

Our research focusses on the development and repurposing of small compound antivirals, including non-nucleoside polymerase inhibitors, TLR7 agonists, and nucleoside analogues for the treatment of infections against positive sense RNA viruses, including norovirus, hepatitis C virus, feline calicivirus and hepatitis E virus. We explore our best antiviral candidates through high-throughput and *in silco* screening, using our cutting-edge enzyme, viral replicons and infection cell-culture based systems.

BIOGRAPHY

After undergraduate studies in Biotechnology from King's College London, Peter completed a PhD at University College, London in bacterial microbiology and molecular biology. In 1996, he started a period of Postdoctoral research at Macquarie University, Sydney, as a recipient of a Royal Society Fellowship to study antibiotic resistance in Gram-negative bacteria. Then in 1998, he joined the Virology Division, Prince of Wales Hospital as Hepatitis Group Leader until 2002, and worked on both HCV and norovirus. In January 2003, he was appointed an academic at UNSW, Sydney and established a molecular microbiology research group and laboratory within the School of Biotechnology and Biomolecular Sciences (BABS). Peter is also the course Coordinator for the third-year course Viruses

and Disease (MICR3061), and he lectures and tutors 2nd and 3rd year science students, as well as medical students at UNSW.



Viral Discovery in Ancient Fish



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Multiple projects are available. These projects provide research training in bacterial pathogenesis, host response to infection, mucosal immunology, bacterial genome, metagenomic analysis, molecular diagnosis of bacterial infection, antibiotics, vaccines for mucosal associated bacteria, or cancer immunotherapy-associated gut microbes.

PROJECTS ON CAMPYLOBACTER CONCISUS AND AEROMONAS SPECIES

Campylobacter concisus is an oral bacterium that may cause enteric diseases. We found that *C. concisus* strains carrie pICON plasmid and pSma1 plasmid are associated with severe Crohn's disease and ulcerative colitis (two major forms of inflammatory bowel disease). *Aeromonas* species are important pathogens of fish, they also cause several human diseases. We recently found that *Aeromonas* species are the third most common enteric pathogens in Australia. Research projects on *Campylobacter* and *Aeromonas* species examine bacterial genomes, virulence factors, bacterial interaction with host immune system, and molecular diagnostic methods.

PROJECTS ON PRECISION ANTIBIOTICS AND VACCINES

Research projects in antibiotic area aim to develop precision antibiotics to specifically eradicate or inhibit individual bacterial species without affecting the balance of gut microbiota. Precision antibiotics may also be used to treat antibiotic resistant pathogenic bacterial species. We are also interested in identifying bacterial components that can be used as vaccines to control mucosaassociated bacterial pathogens.

PROJECTS ON CANCER IMMUNOTHERAPY-ASSOCIATED GUT MICROBES

Blockade of immune checkpoint proteins is a type of cancer therapy. Recent studies found that some bacterial species in the gastrointestinal tract may affect the efficacy of immune checkpoint blockade therapy. Projects in this area investigate the mechanisms by which gut bacterial species affecting cancer immunotherapy, aiming to provide additional strategies to improve cancer immunotherapy efficacy.

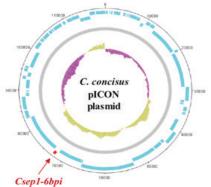


Figure 1. Circularised diagram of the pICON plasmid in *C. concisus* strain P2CDO4. (doi:10.1038/s41426-018-0065-6)

RESEARCH FOCUS

Campylobacter, Aeromonas and other mucosa-associated bacteria, chronic inflammatory diseases, cancer immunotherapy-associated microbes.



BIOGRAPHY

Associate Professor Li Zhang received MBBS degree from Fudan University in Shanghai, China and PhD degree from the University of Cambridge in the UK. A/Prof Zhang was a clinician at the China-Japan Friendship Hospital in Beijing prior to her PhD study. She worked as a postdoctoral fellow at the Institute of Molecular and Cell Biology in National University of Singapore before joining the University of New South Wales.

RESEARCH

A/Prof Zhang's group investigates bacterial species that cause or prevent inflammatory diseases and cancers of the gastrointestinal tract. They study bacterial genomes, bacterial virulence factors, interactions of bacterial pathogens and gut microbiome with the immune system, and novel methods to modify gut and oral microbiome. A/Prof Zhang's earlier research was on autoimmune diseases.

A/Prof Li Zhang is a pioneering researcher in the field of human hosted *Campylobacter* species and inflammatory bowel disease (IBD). She hypothesized that some strains of *C. concisus*, a bacterium that usually colonises the human oral cavity, have enteric pathogenicity and are the initiator of a subgroup of human IBD. Research in A/Prof Zhang's group has provided critical information to this research field, including disease associations, *C. concisus* natural colonisation site, *C. concisus* metabolic pathways and *C. concisus* genomic features associated with severe IBD.

Aeromonas species are important pathogens of fish and emerging human pathogens. A/Prof Zhang group recently reported that Aeromonas species are the third most common gastrointestinal bacterial pathogens in Australia, following Campylobacter and Salmonella species. They are currently examining the pathogenic mechanisms of Aeromonas species in causing human diseases.

Evidence shows that bacterial species in the gastrointestinal interesting tract affecting cancer immunotherapy A/Prof Zhang group investigates the mechanisms by which gut bacterial species affecting cancer immunotherapy. They also develop precision antibiotics to modify gut microbiota, aiming to provide additional strategies to improve cancer immunotherapy efficacy.

For other details about research in A/Prof Zhang group and their publications, please visit:

- research.unsw.edu.au/people/associate-professor-lizhang
- scholar.google.com.au/ citations?user=gddDrbMAAAAJ&hl=en

RESEARCH SUPERVISORS -

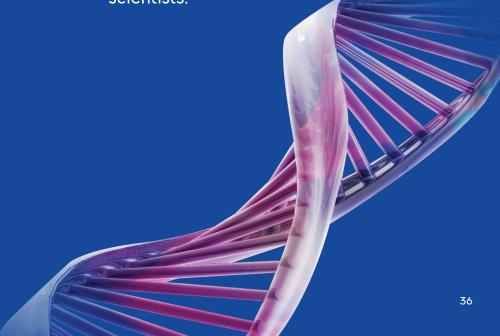
MOLECULAR AND CELL BIOLOGY

CLUSTER STRENGTHS:

- » Metabolism and Metabolic Disorders
- » Structural and Synthetic Biology

Since traditional biology focuses on living organisms as a whole, Molecular and Cell Biology explores the components and interactions that make up a cell. This gives us a deeper understanding of cell function and why diseases and disorders happen on a molecular level.

Molecular and Cell Biology has been pivotal in a wide range of fields and revolutionised the ability to manipulate cells and tissues for medical and therapeutic purposes such as vaccinations. Other developments have included DNA fingerprinting in forensics and pioneering crop modifications in agriculture. Our research centres on the areas of Synthetic Biology and Metabolism and Molecular Cell Biology. We incorporate molecular genetics, stem cell biology, microscopy, computer science and epidemiology to answer unsolved biological questions and train the next generation of life scientists.





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RESEARCH FOCUS

- Evolutionary Biophysics
- Synthetic Biology
- Super resolution microscopy



My research group currently focuses on three streams of research:

- Molecular evolution of the bacterial flagellar motor

 how did this motor arise and how can we use synthetic biology to create new motors?
- 2. Controlling fluid flows with bacteria how can we engineer the motor to be light responsive and to propel mixing in microfluidics?
- 3. Building new cells out of DNA, lipids and proteins – how can we build synthetic, light-controllable cells using engineering principles and biological components we buy off the shelf?

For more info about Matt and all the people in the lab please visit the lab <u>website</u>.



PROJECT 1 - ANCESTRAL RECONSTRUCTION OF THE BACTERIAL FLAGELLAR ROTOR

We recreate microbial 'Jurassic Parks' by resurrecting ancient flagellar motor componentry in contemporary hosts and measuring how well they work. This allows us to create ancient motors that have never existed in the present day to synthesise and evolve new motors as

well as to learn about the process of evolution. We have examine in detail reconstructions of the stators that power the motor and now seek to examine how the rotor has evolved and can be engineered for new applications.

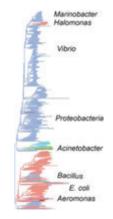


PROJECT 2 - EVOLUTION ACROSS INTERFACES

In this project we explore the directed evolution of the flagellar motor in the lab by evolving it to swim under different energy sources and selecting for motility. Recent work in antibiotic resistance (eg by Michael Baym) has shown that the resistance of antibiotics occurs in lockstep when progressing through 10-fold increases in antibiotics. We aim to explore how motility evolves across interfaces, when a bacterium faces a change in environment between, for example, H+ and Na+ environments, and how the bacteria adapts to dwindling nutrient across this interface. This project has scope for designing and building custom tanks to optimise bacterial evolution using 3D printing and prototyping, as well as investigating microbiology and bacterial motility in multiple dimensions using layered swim devices.

PROJECT 3 - ORIGINS OF MOTILITY

The evolutionary origins of the bacterial flagellum have been a subject of scientific and public controversy – how can evolution produce such a complex system? We believe we can make progress on the issue by updating old phylogenetic work with new datasets and improved models, and combining this with experimental evolution work being done in our labs. The project will be to assemble a well- organized database of flagellar proteins and explore sequenced bacterial genomes with genome browsers and similarity



searches. The student will identify flagellar proteins and their evolutionary relatives, including recording their position in the genome. The student will also plan and conduct phylogenetic analyses, and then use synthetic biology to recreate these ancestors in a contemporary microbial 'Jurassic Park'.

PROJECT 4 - REGULATION OF MEMBRANE PROTEIN INSERTION IN ARTIFICIAL BILAYERS USING DNA ORIGAMI

Our droplet hydrogel bilayer system is an artificial bilayer system for interrogating membrane proteins, but it also allows us to explore new forms of synthetic biology where we can add individual protein function to a droplet, such as touch sensitivity or light sensitivity. Using a DNA origami nanostructures we can protect and controllably release our blocking DNA structures to direct the fusion of liposomes and control which reactions take place where in these droplets. This allows us to trigger functionality, on demand, using light and electrical signals. This project involves in vitro synthetic biology, DNA and lipid nanotechnologies and microscopy.

PROJECT 5 - APPLICATIONS OF FLAGELLAR MOTOR TO FLUID FLOWS

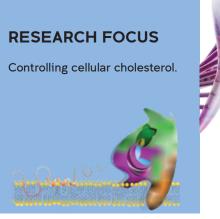
We utilise the high efficiency and self assembly of the flagellar motor to drive rotation of cells on patterned surfaces to control mixing and fluid flows in microfluidics. We have projects involving designing and building new devices to apply the flagellar motor onto other things. This would suit someone with an interest in DIY/maker culture.







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RESEARCH "I <3 Cholesterol, but my <3 doesn't"

However, there's much much more to cholesterol than that. We study how we and our cells balance this Jekyll/Hyde molecule.

Cholesterol is notorious in human health and disease. It is both vital and lethal, depending on its levels, which are determined by several factors, including cholesterol synthesis. For the past decade, my lab has focused on investigating the control of cholesterol synthesis. We have made major progress, uncovering novel modes of regulation of enzymes beyond the best known and most intensively studied enzyme (HMGCR, target of the statin class of drugs). Notably, we discovered an important control point later in the pathway (squalene monooxygenase or epoxidase, SQLE), which has recently become the subject of intense interest as an oncogene and therapeutic target in several cancers.

RESEARCH GOALS

- To discover new factors in achieving cholesterol balance in cells
- To identify links between cholesterol and cancer

SPECIFIC PROJECTS

Cholesterol is a vital and versatile molecule that has become a byword for heart disease risk. In fact, the cells in our body actually need cholesterol, and too little results in devastating developmental disorders. However, too much can contribute to several diseases, including atherosclerosis and cancer. Our bodies have therefore engineered an elaborate system for keeping the cholesterol content of our cells tightly controlled. The overall goal of our research is to understand more about how our cells control cholesterol levels.

PROJECT 1: New factors in achieving cholesterol balance

An imbalance of cholesterol plays a role in numerous diseases. Therefore, knowing precisely how cells regulate their cholesterol levels is central to understanding the development of these diseases, and to identify possible new treatments. Only one of the 20+ enzymes involved in cholesterol biosynthesis is targeted clinically (by statins). The statin class of drugs, worth >\$30 billion a year, inhibit a very early step in cholesterol synthesis and have been effective in treating heart disease, but are not without their side effects. Very little attention has been paid to later steps in the pathway. This project will investigate the regulation of new control points in cholesterol synthesis, which have been largely overlooked in the past. The statin class of drugs, worth >\$30 billion a year, inhibit a very early step in cholesterol synthesis and have been effective in treating heart disease, but are not without their side effects. Very little attention has been paid to later steps in the pathway.

This project will investigate the regulation of new control points in cholesterol synthesis, which have been largely overlooked in the past. in the pathway. This project will investigate the regulation of new control points in cholesterol synthesis, which have been largely overlooked in the past.

PROJECT 2: Cholesterol and cancer

Cancer is a disease characterised by increased cellular replication and spread beyond the normal location in the body. A hallmark feature of cancer cells is their abnormal metabolism compared to normal cells. Notably, cells need cholesterol to grow and proliferate and mechanisms to accumulate cholesterol are far more common in cancer cells. Our lab discovered a connection between a major player involved in maintaining cholesterol balance in animal cells and a key proliferative pathway that is overactive in many cancers, including prostate cancer. This project investigates novel ways to modulate and decrease cellular cholesterol levels, which may inform the development of new anti-cancer therapies.

METHODS ROUTINELY USED IN THE LAB

Mammalian cell culture, recombinant DNA techniques (cloning and mutagenesis), fluorescence microscopy, real-time PCR, gene/siRNA transfection, luciferase reporter assays, SDS-PAGE, Western blotting, and mass spectrometry.

SUPERVISION OPPORTUNITIES/AREAS

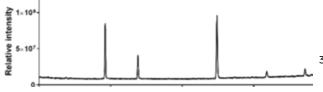
Our lab provides a nurturing, supportive and stimulating research environment for students.

I have supervised 26 Honours students and all have received first class honours; 15 have gone on to do a PhD with me. Seven of my Honours students have received University medals (top of their cohort).

I have supervised 10 PhD students and all obtained excellent post-doc positions in Australia or overseas after completion.

I believe that a key part of my role as an academic is to mentor the next generation in scientific publishing, so I have an established record of converting student projects into publications. Many of my Honours students have had work from their Honours year published, and my PhD students publish at least 5 papers on average during their time with me.

Contact Andrew about research supervision opportunities: aj.brown@unsw.edu.au





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My lab is interested in understanding the relationship between obesity and cancer, and whether we can target obesity to improve outcomes for patients with cancer.



Dr Byrne completed her PhD at the Children's Cancer Institute, UNSW, and trained as a postdoc at the University of Virginia (USA). Her laboratory in the School of BABS focuses on developing new drugs to treat cancer.

Research Project:

Developing new treatments for obesity-related cancers

Obesity is linked to the development of 13 different types of cancer. Obese individuals often have increased levels of circulating pro-tumour growth factors such as insulin, and tumour-promoting microenvironments of inflammation, fat accumulation and oxidative stress in organs such as the liver. These obesity-related factors promote cancer development and progression, as well as cancer recurrence and drug resistance. However, obesity and its related protumour factors are not currently considered when treating cancer

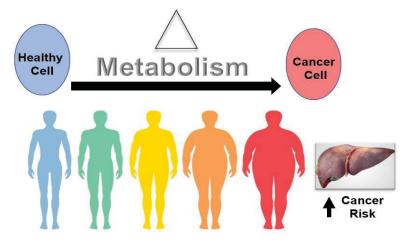
My laboratory is investigating whether targeting obesity, using novel mitochondrial uncoupler molecules such as BAM15 (Kenwood et al. 2014), improves the anti-cancer response to cancer therapies. Mitochondrial uncouplers are small molecules that make mitochondria burn (oxidise) more nutrients, such as fats, without producing ATP. Thereby they 'uncouple' nutrient oxidation from ATP synthesis. These molecules have shown great promise for the treatment of obesity. However, we will test whether targeting obesity with these uncouplers enhances the anti-tumour effects of cancer therapies using in vitro and in vivo models. These experiments will help us determine whether uncouplers may be used as novel and effective treatments for obesity-related cancers in the future.

Recent publications:

- Shrestha R, ..., Byrne FL. Complete inhibition of liver acetyl-CoA carboxylase activity is required to exacerbate liver tumorigenesis in mice treated with diethylnitrosamine. Cancer Metab. 2024 Nov 13;12(1):34. doi: 10.1186/s40170-024-00363-1. PMID: 39538356
- Cheng Y, ..., Byrne FL, Kumar N. Design, synthesis and biological evaluation of glucose metabolism inhibitors as anticancer agents. *Bioorg Chem.* 2024 Oct;151:107665. doi: 10.1016/j.bioorg.2024.107665. Epub 2024 Jul 25.
- Hargett S, ..., Byrne FL. Bile acids mediate fructoseassociated liver tumour growth in mice. Biochim Biophys Acta Mol Basis Dis. 2024 Mar;1870(3):167029. doi: 10.1016/j.bbadis.2024.167029. Epub 2024 Feb 6. PMID: 38325224.

Publications related to mitochondrial uncouplers:

- Shrestha, R., Johnson, E., Byrne, F.L. (2021). Exploring the therapeutic potential of mitochondrial uncouplers in cancer. *Mol Metab.* 2021;51:101222. doi:10.1016/j. molmet.2021.101222
- Kenwood, B. M., ..., Byrne, F. L., ...Hoehn, K. L. (2013). Identification of a novel mitochondrial uncoupler that does not depolarize the plasma membrane. *Molecular metabolism*, 3(2), 114–123. https://doi.org/10.1016/j.molmet.2013.11.005
- Alexopoulos, S. J., Chen, S. Y., Brandon, A. E., Salamoun, J. M., Byrne, F. L., ... Hoehn, K. L. (2020). Mitochondrial uncoupler BAM15 reverses dietinduced obesity and insulin resistance in mice. Nature communications, 11(1), 2397. https://doi.org/10.1038/s41467-020-16298-2





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Immune response to infection, injury and cancer.



RESEARCH PROGRAM

Our main goal is to understand and control inflammation.

During infection, inflammation enables immune cells to locate and eliminate invading pathogens. However, chronic inflammation can cause tissue damage in autoimmune diseases and contribute to cancer development. Our lab seeks to understand how immune cells respond to different types of inflammation. This knowledge will allow us to manipulate inflammatory responses and convert detrimental immune activity into beneficial outcomes.

We are currently recruiting Honours and PhD students to work on projects that aim to:

- Utilize single-cell sequencing approaches to investigate immune responses to infection and cancer immunotherapy.
- Uncover the role of immune cells in mediating wound repair using intravital microscopy and flow cytometric approaches.

Both projects require strong coding skills and involve computational biology. A solid understanding of immunology and cell biology would be beneficial.

For international students, we are only able to accept PhD candidates with scholarships.

Contact Tatyana for research project opportunities: t.chtanova@unsw.edu.au

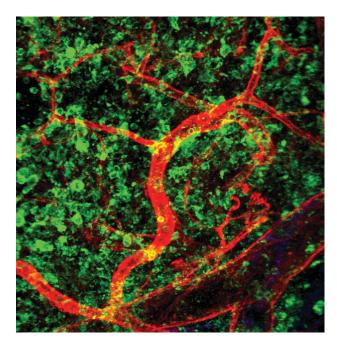
BIOGRAPHY

Associate Professor Tatyana Chtanova is the head of the Innate and Tumour Immunology Laboratory at the School of Biotechnology and Biomolecular Sciences, UNSW Sydney.

After completing her undergraduate studies at the University of New South Wales, Tatyana earned her PhD in 2005 for her research on gene expression signatures in novel T cell subsets, conducted at the Garvan Institute.

Following her PhD, she received a Human Frontier Science Program Fellowship to undertake postdoctoral training at the University of California, Berkeley. During this time, she developed expertise in intravital microscopy and applied it to uncover a distinctive immune response to inflammation—neutrophil swarming—and a novel mechanism of immune evasion by pathogens.

Tatyana's research focuses on developing advanced imaging techniques such as two-photon microscopy and in situ photoconversion to investigate fundamental immunological processes, including infection, wound repair, and cancer. Her overarching goal is to harness inflammation to develop new immunotherapies for cancer and enhance wound healing.





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Structural Biology, synthetic biology, protein structure and function, biophysics, protein chemistry, X-ray crystallography, cryo-electron microscopy.

Our research focuses on understanding how proteins work at the atomic level. Proteins are nature's choice for making cellular machines. Each protein machine is composed of well-ordered structural domains that are linked together to create a dynamic, functioning system. By mapping the structures of a protein in action, we can create a series of snapshots that reveal how the protein works in the cell. We also track protein evolution across large timescales to gain a deeper understanding in the context an organism. While traditionally, we have studied natural protein systems in order to understand function, a new challenge is to use our knowledge to design and create new protein machines using the tools of synthetic biology.

PROJECT 1

How do protein motors work? Nature has evolved spectacular protein motors such as myosin and kinesin that can "walk" down protein tracks (actin filaments and microtubules, respectively). Although these motor proteins have been studied for decades, producing a plethora of atomic structures and detailed mutagenic studies, we still have no idea how these proteins harness chemical energy (ATPase) to produce motion. A novel way to explore this question is to use a synthetic biology approach: take existing protein modules of known function and link them together to create artificial protein motors. Our lab is part of an international team that is striving to achieve this goal. We have developed a successful strategy to link functional

PROJECT 2 (jointly supervised with Dr Kate Michie)
How does a eukaryotic cell control the shape of the
plasma membrane and associated vesicles? Underlying
the eukaryotic plasma membrane is a layer of actin
filaments called the cell cortex or cortical cytoskeleton.
The protein ezrin (and its paralogues, the ERMs) couple

membranes to cortical actin filaments. Ezrin is responsible for the maintenance of surface structures (such as microvilli) and the invagination of the plasma membrane during processes such as phagocytosis. Our key question is how does ezrin do this? We have determined crystal structures of ezrin in two states,

modules together to make an artificial motor protein that will

"walk" along a DNA nanotube

track. This project will explore

the same components as our

a new motor design using

current motor.

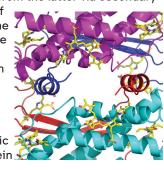


however, we still lack a structure of the active, membrane bound form of ezrin. Preliminary cryo electron microscopy studies show that ezrin alone can deform membrane vesicles and cluster them together. The aim of this project is to determine how ezrin achieves this, with the ultimate aim of obtaining the structure of ezrin bound to a membrane, and, finally, to actin filaments.

PROJECT 3

How do light harvesting proteins capture sunlight and transmit the energy so as to power photosynthesis? Aquatic organisms have evolved elaborate light harvesting antenna systems where proteins control the capture and transfer of energy between chromophore molecules. During evolution, at least five distinct light harvesting antenna systems have been discovered. Our research focuses on the light harvesting antennae of two classes of algae: the cryptophytes and red algae, where the former evolved from the latter via secondary

endosymbiosis. The aim of this project is to explore the dramatic changes that have occurred in the antenna complexes during evolution using synthetic biology approaches. By creating protein chimeras, we will explore how changes in sequence result in dramatic structural changes in protein complexes.



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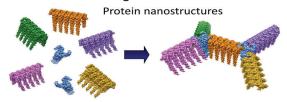
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Synthetic biology and bioengineering of protein biomaterials.

RESEARCH PROGRAM

The intricate and ordered complexes that proteins adopt in nature is central to many biological processes, ranging from cellular scaffolding provided by cytoskeletal proteins to the encapsulation of nucleic acids in viral capsids. Exploiting this remarkable fidelity and precision in self-assembly is highly attractive for the fabrication of structurally defined materials with nanometer dimensions. Advances in the computational prediction of protein folding have enabled the design of proteins that self-assemble into complex yet predictable shapes.

The research program in my group applies synthetic biology for the engineering of proteins into structured and functional biomaterials. Central to this approach is the creation of standardised protein building blocks that can be assembled into geometrically-defined structures of controllable size and shape. This ability to design protein nanostructures with atomic-level accuracy opens new possibilities in biomaterials. Applications of these protein-based biomaterials include the creation of electrically conductive protein nanowires for biosensors and enzyme catalysis, the engineering of metabolic pathways, and the fabrication of tissue scaffolds for regenerative medicine.



SPECIFIC PROJECTS

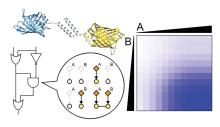
Project 1: Conductive protein nanowires for bioelectronics and biosensors

The recent discovery of electrically conductive proteinbased nanowires produced by bacteria has potential applications in the development of bioelectronics and biosensors. Exploiting this conductivity and the ability of proteins to self-assemble into complex structures may facilitate the fabrication of structured nanoscale devices that can directly interface with biological systems (e.g., enzymes or living cells). This project will create novel protein nanowires by alignment of redoxactive proteins on filamentous scaffolds. Subsequently, the protein nanowires will be used to mediate the transmission of electrons for novel electrical devices such as biosensors or for direct communication with living cells.

Conductive protein nanowires

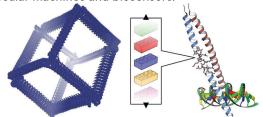
Project 2: Design of synthetic transcriptional factors

An aim of synthetic biology is to engineer useful genetic systems inside living cells - for example, to make cells produce drugs or detect changes in the environment. The challenge is: can synthetic genetic circuits interfere with the rest of the cell? In this project, we will build synthetic transcription factors that can be used to regulate synthetic genetic circuits. Conversely, synthetic transcription factors can also be used to modulate natural genes in a controllable manner. The applications of synthetic transcription factors extend from the design of synthetic living systems to targeted gene/protein therapies for genetic diseases.



Project 3: Self-assembling biomaterials for nanotechnology

The fabrication of nanoscale devices requires architectural templates upon which to position functional molecules in complex arrangements. Protein and DNA are attractive templates for nanofabrication due to their inherent self-assembly and molecular recognition capabilities. This project will engineer a new class of biotemplates that use DNA origami to link filamentous proteins into three-dimensional templates of controllable size and symmetry. Subsequently, these novel biotemplates will serve as a foundation upon which to build functional nanodevices including molecular machines and biosensors.



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RESEARCH FOCUS

- Immunology
- •2-photon microscopy
- ·CRISPR Gene editing
- Mouse models
- Mouse genome editing
- Flow cytometry
- Molecular biology
- ·Human genetics •RNA sequencing

nature

immuno

TNFAIP3 vari

·Whole genome sequencing

IMPACT:

Nat Immunol, 2019

Oct;20(10):1299-1310

The Transplantation Immunology Laboratory GARVAN INSTITUE OF MEDICAL RESEARCH

*Why do some people develop immune diseases and others not?

*Why does COVID19 have worse symptoms in particular people?

*Why are immune cancer therapies more effective in some people?

We are addressing these key questions by exploring how human genetic variation in an immune master controller gene called TNFAIP3 effects us in health & disease.

Projects suitable for Honours, Masters and PhD studies are available.

CONTACT: Prof Shane T. Grey: s.grey@ unsw.edu.au

IMAGE: Pancreatic Islets, the

target in T1D

EXCITING SCIENCE THROUGH COLLABORATION:













FUNDING:







RESEARCH SUPPORT







Professor Kyle Hoehn
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Mechanisms linking altered nutrient metabolism to ageing, obesity, and diabetes. Suitable for students who have majored in Molecular Biology.

Kyle earned a PhD in Biochemistry from Colorado State University (2000-2005) and completed postdoctoral training at the Garvan Institute of Medical Research in Sydney (2005-2009). He then established his independent lab at the University of Virginia as an Assistant Professor of Pharmacology in 2009 and moved his lab to UNSW School of BABS in 2014. Kyle co-founded Continuum Biosciences in 2017 and Uncoupler Biosciences in 2023.

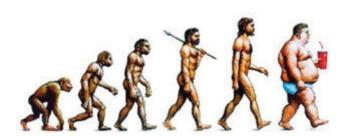
Research in the Hoehn lab aims to better understand metabolic processes in normal conditions, ageing, and metabolic disease states. Projects include:

Project 1: Developing next-generation mitochondrial uncouplers

Mitochondrial uncoupler molecules are anti-oxidant and boost metabolic rate without affecting food intake. Mitochondrial uncouplers have potential uses for obesity, diabetes, anti-ageing, and other disorders.

The Hoehn lab discovered the furazanopyrazine class of mitochondrial uncouplers. The first molecule discovered was named BAM15 after undergraduate student Beverley A. Murrow. BAM15 is now sold commercially by many vendors and is used widely in medical research (give it a google). BAM15 is a great starting point for drug development and the lab is continually developing new molecules to improve drug-like properties that are better suited for human use.

Students that join this project will test our molecules in cells and mice for pharmacokinetics, safety, and efficacy to reverse obesity, fatty liver disease, or agerelated decline.



Project 2: Determining the effect of Acetyl-CoA Carboxylase activity on protein acylation

Acetyl-coenzyme A carboxylase (ACC) enzymes are rate limiting for fat synthesis. ACC enzymes consume acetyl-CoA to produce malonyl-CoA, which is the major building block for fat synthesis. Recently we have identified that ACC enzymes also regulate post-translational modification of hundreds of proteins by limiting the availability of acetyl-CoA and malonyl-CoA for protein acetylation and malonylation, respectively. We are performing proteomics studies to determine which proteins are sensitive to ACC-dependent acetylation and malonylation.

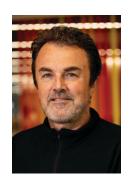
Student projects will characterise the functional consequences of ACC-dependent acetylation or malonylation of key metabolic proteins. Students will use gene editing to block or add acetyl/malonyl marks and then determine the effect on enzyme activity.

RELEVANT PUBLICATIONS:

2023 – Chen SY, Beretta M, Olzomer EM, Shah DP, Wong DYH, Alexopoulos SJ, Aleksovska I, Salamoun JM, Garcia CJ, Cochran BJ, Rye KA, Smith GC, Byrne FL, Morris MJ, Santos WL, Cantley J, <u>Hoehn KL</u>. Targeting negative energy balance with calorie restriction and mitochondrial uncoupling in db/db mice. <u>Mol Metab</u>. Mar;69:101684. doi: 10.1016/j.molmet.2023.101684.

2021 - Chen SY, Beretta M, Alexopoulos SJ, Shah DP, Olzomer EM, Hargett SR, Childress ES, Salamoun JM, Aleksovska I, Roseblade A, Cranfield C, Rawling T, Quinlan KGR, Morris MJ, Tucker SP, Santos WL, and <u>Hoehn KL</u>. Mitochondrial uncoupler SHC517 reverses obesity in mice without affecting food intake. <u>Metabolism</u>. Apr;117:154724. doi: 10.1016/j. metabol.2021.154724.

2020 – Alexopoulos, A.J.; Chen, S-Y.; Brandon, A.E.; Salamoun, J.; Garcia, C.J.; Beretta, M.; Olzomer, E.; Byrne, F.B.; Shah, D.; Lawrence, R.; Carrive, P.; Tucker, S.P.; Cooney, G.J.; Santos, WL; <u>Hoehn KL</u>. Mitochondrial uncoupler BAM15 reverses diet-induced obesity and insulin resistance in mice. <u>Nature Communications</u>. May 14;11(1):2397. doi: 10.1038/s41467-020-16298-2.



Associate Professor Christopher Marquis

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Bioprocessing Bioproduction



RESEARCH PROGRAM

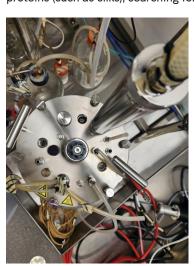
A/Prof Marquis trained as a biochemical engineer and is a current academic member of BABS. In addition, A/Prof Marquis is the Director of the Recombinant Products Facility (www.proteins.unsw.edu.au), which he established in 2008. The RPF houses research infrastructure for fermentation, cell culture, mid-stream and downstream processing for protein production, purification and characterisation. The facility also provides expertise in developing and optimising bioprocesses. The primary role of the Facility is to provide research support to the UNSW research community, however the facility provides contract services for the wider research community and also industry. The RPF has fermentation capacity to 20L and a variety of mid and downstream processing capabilities for the production and purification

of proteins for research. In addition to running contract projects the facility has a number of internal initiatives to investigate and optimise bioprocesses, including the use of microbioreactor systems for process optimisation.



A/Prof. Marquis has current interdisciplinary

research projects across from basic since through to industry-linked projects. Current projects include investigating the role of tyrosine in phase-separating proteins (such as silks), searching for and developing



improved versions of the enzyme gamma glutamyltransferase for dipeptide bioproduction and investigating circular RNAs that encode protein production. New projects include those with industrylinks; BondiBio and Smart MCs. Here is a summary of current projects available for students:

MARQUIS LAB PROJECTS FOR STUDENTS

PROTEIN AND NUCLEIC ACID PROJECTS

- SEARCHING FOR NOVEL ENZYMES FOR DIPEPTIDE SYNTHESIS (Hons/MSc); Gammaglutamyl transferase is a ubiquitous enzyme and is found to have use in the production of the dipeptide gamma glutamyl cysteine. Currently, the enzyme is sourced from native mammalian tissue. This project will explore alternative native and recombinant methods to generate active enzymes for improved industrial application of this enzyme.
- 2. OPTIMISING scPLASMID DNA PRODUCTION (Hons/MSc); Plasmid DNA is routinely produced in laboratories worldwide for a range of fundamental and applied purposes. For application in a therapeutic context, plasmid yield and quality is critical and not commonly addressed in most labs. This project will involve a systematic evaluation of variables that generates high yields of supercoiled plasmid DNA for downstream applications such as direct gene delivery as well as templates for in vitro transcription to produce mRNA.

INDUSTRY-LINKED PROJECTS

- I. NOVEL MICROCARRIERS FOR MAMMALIAN CELL BIOPROCESSING (Hons/MSc/PhD); Cultured mammalian cells produce valuable biopharmaceuticals (including viral gene therapy vectors) and potentially provide tissues for autologous transplantation therapies. This project will focus on the design and use of novel microcarriers for the production valuable bioproducts (proteins and viruses), using attachment-dependent cell lines. This project will work closely with Professor Kris Killian (Chemistry) and SmartMCs and is supported by an ARC Linkage project grant.
- 2. CYANOBACTERIA AS A PLATFORM FOR MAKING RECOMBINANT ANTIBODIES (Hons); This project is in collaboration with locally-resident **Bondi Bio** project will investigate the use of cyanobacteria to produce monoclonal antibodies. The proteins are effectively mass-produced currently in mammalian cell culturebased processes, however the cost of production is still significant. Cyanobacteria are phototrophs that can grow on simple media in photobioreactors at large-scale, presenting potentially a lower cost production option for these valuable proteins. This project will look at assessing important variables to improve productivity, working closely with Bondi Bio staff.



Dr Kate Michie Senior Research Associate and Lecturer

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RESEARCH FOCUS

Structural biology of protein machineries.



RESEARCH AREAS

We want to understand how biology uses proteins to control the shape of membranes. In particular we focus on the proteins essential to the processes of cell division and building cell surface structures such as villi and vesicles. The proteins involved in these processes interact with themselves and with the membrane to bend and shape it, and to tether other proteins to it. The types of proteins involved arise in all the domains of life— archaea, bacteria and eukaryotes, and, maybe not surprisingly, are often carried out by related proteins.

We focus on understanding structurally the proteins that carry out these tasks. We use X-ray crystallography, cryo-electron microscopy and a range of protein and biophysical experiments to probe our targets.

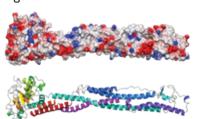


Figure 1: Electrostatic potential and crystal structure of LeoA- a bacterial dynamin-like protein.

CURRENT RESEARCH ACTIVITIES

- · Dynamin-like Proteins in Bacteria
- Tubulin-like proteins in Bacteria and Archaea
- Ezrin/Moesin and Merlin and the membrane in humans (Jointly supervised with Prof Curmi)

Dr Michie also collaborates other researchers. If you are interested in structural biology, introduce yourself to explore potential paths forward.

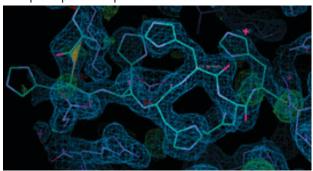


Figure 3: Electron density and protein model showing a his-tag on a recombinant protein structure

BIOGRAPHY

Dr Kate Michie is an Adjunct Lecturer (BABS) and Senior Research Associate in the Mark Wainwright Analytical Centre. Dr Michie completed her PhD at Sydney University (2004). She was an International L'Oreal UNESCO Fellow (2005) and Marie Curie Fellow (2006-2007) at the Medical Research Council Laboratory of Molecular Biology (LMB) in Cambridge UK with Dr Jan Löwe, and a Research Associate at St John's College. She was made an Investigator Scientist at the LMB in 2008 working on the structural biology of bacterial cytoskeletal proteins. Returning to Australia in 2010, Dr Michie worked on heart proteins with Professors Guss and Trewhella (Syd Uni). She moved to UNSW in 2016 to work with Professor Paul Curmi on Ezrin (a membrane shaping cytoskeletal protein) and light-harvesting complexes. In 2019 she was employed to set up and run the Structural Biology Facility within the Mark Wainwright Analytical Centre.

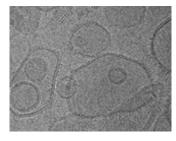


Figure 2: Cryo EM image of Ezrin binding on the surface in regular arrays to lipid vesicles.

SELECT PUBLICATIONS

- Scaffolding proteins guide the evolution of algal light harvesting antennas. HW Rathbone, KA Michie, MJ Landsberg, BR Green, PMG Curmi. Nature communications 12 (1), 1-9, 2021
- Two sides of the coin: ezrin/radixin/moesin and merlin control membrane structure and contact inhibition. KA Michie, A Bermeister, NO Robertson, SC Goodchild, PMG Curmi. *International journal of* molecular sciences 20 (8), 1996. 2019
- CetZ tubulin-like proteins control archaeal cell shape. IG Duggin et al. Nature 519 (7543), 362-365. 2015
- LeoA, B and C from enterotoxigenic Escherichia coli (ETEC) are bacterial dynamins. KA Michie, A Boysen, HH Low, J Møller-Jensen, J Löwe. *PloS* one 9 (9), e107211. 2014



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expenditure in adipose tissue.

Regulation of energy

We study mammalian metabolism and gene regulation, with the aim of identifying biological pathways to target for anti-obesity therapies. White adipose tissue can be converted to 'beige' adipose tissue, which burns energy to produce heat rather than storing energy. We aim to better understand beige adipose tissue so that this knowledge can be harnessed to reverse obesity.

Currently, our collaborative research group includes a lab manager, 7 PhD students and 2 Honours students. We are seeking new HDR students for 2026.

TECHNIQUES

Our projects offer the opportunity to learn a wide variety of molecular biology and cell biology techniques, including chromatin immunoprecipitation (ChIP), western blotting, gel shifts, subcloning and bacterial transformation, site directed mutagenesis, CRISPR/ Cas9 genome editing, PCR and real-time PCR, next-generation sequencing technologies (RNA-seq and ChIP-seq), tissue culture, transient and stable transfections of mammalian cells, reporter gene assays and flow cytometry.

BIOGRAPHY

Kate is a Scientia Associate Professor in the School of Biotechnology and Biomolecular Sciences (BABS). She runs a research group interested in gene regulation, with a particular focus on understanding the communication between immune cells and fat cells within adipose tissue to uncover new therapeutic targets for obesity. Kate completed her Bsc (Hons) (Advanced) at the University of Sydney. She went to overseas to work as a Research Assistant at the University of Cambridge for a year before returning to the University of Sydney to complete her PhD in transcription factor biology. Kate moved to the

Children's Hospital at Westmead as a post-doc, where she studied the effects of the human ACTN3 gene polymorphism on skeletal muscle performance and metabolism. She continued her post-doctoral training at the University of Cambridge (2008-2010), researching pluripotency of embryonic stem cells and targeted differentiation of these cells towards skeletal muscle satellite cells. She then returned to the University of Sydney to lead a program of research into the human ACTN3 gene polymorphism, with the aims of uncovering mechanisms behind changes to muscle function and metabolism in individuals homozygous for this polymorphism, and translating findings from model organisms to humans. In 2014 Kate joined UNSW, to study the transcriptional regulation of haematopoiesis and transcription factor mechanisms in addition to continuing an independent research program focussed on skeletal muscle metabolism. She became an independent group leader in 2018. Kate mentors a number of PhD and Honours students.

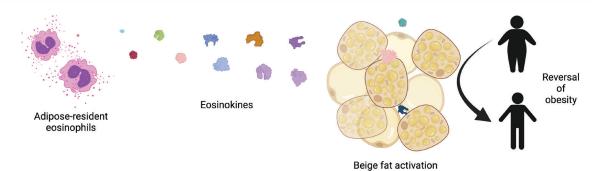
RECENT PUBLICATIONS

'Shades of white: new insights into tissue-resident leukocyte heterogeneity' FEBS J., 2022, 289(2):308-318.

'Eosinophil Function in Adipose Tissue Is Regulated by Krüppel-like Factor 3 (KLF3)' Nature Communications, 2020, 11(1):2922.

'Krüppel-like Factor 3 (KLF3) Suppresses NF-κB-driven Inflammation in Mice.' J Biol Chem., 2020, 295(18):6080-

'EoTHINophils: Eosinophils as Key Players in Adipose Tissue Homeostasis' Clin Exp Pharmacol Physiol, 2020, 47(8):1495-1505.



We propose that eosinophils drive beige fat activation by producing novel signalling proteins that we term 'eosinokines' which we aim to harness to reverse obesity



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RESEARCH FOCUS

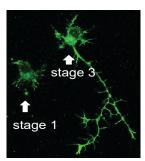
Neurobiology, neuroscience, recognition and cell adhesion in neurons.

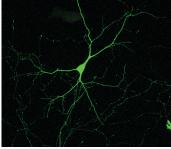


RESEARCH INTERESTS AND CONTRIBUTIONS

In the brain, information is transmitted, processed and memorised by neurons. To perform these functions, neurons must grow and form networks, in which individual neurons are connected to other neurons by specialised contacts called synapses. Neurons use synapses to communicate with other neurons and to process and store information.

The formation and maintenance of the neuronal networks and synapses is regulated by neural cell adhesion molecules expressed at the cell surface of neurons (see our review Sytnyk et al., Trends in Neurosciences, 2017). Our laboratory is interested in understanding the molecular and cellular mechanisms of this regulation and effects of its loss in disease. We also develop new technologies aimed at improving brain performance, enhancing learning and maintaining memory by modulating neural cell adhesion molecules.





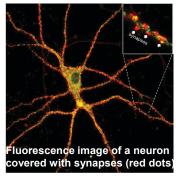
My early work showed that neural cell adhesion molecules are the first proteins accumulating at nascent synaptic contacts between developing neurons and that these proteins stabilize the contacts and induce their transformation into mature synapses by capturing synaptic precursor organelles (Sytnyk et al., Journal of Cell Biology, 2002, featured on the cover page). We then found that neural cell adhesion molecules regulate the key processes involved in neuronal growth and synapse formation including intracellular signalling (Leshchyns'ka et al., Journal of Cell Biology, 2003; Bodrikov et al., Journal of Cell Biology, 2005, 2008; Sheng et al., Journal of Neuroscience, 2015), the assembly of the cytoskeleton (Puchkov et al., Cerebral Cortex, 2011; Li et al., Journal of Neuroscience, 2013) and polarised intracellular transport (Chernyshova et al., Journal of Neuroscience, 2011). We demonstrated that neural cell adhesion molecules modulate the assembly and maturation of the neurotransmitter-releasing machinery in axons (Leshchyns'ka et al., Neuron, 2006; Shetty et al.,

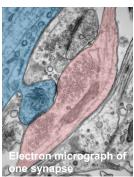
Journal of Neuroscience, 2013) and neurotransmitterdetecting machinery in dendrites of neurons (Sytnyk et al., Journal of Cell Biology, 2006; Sheng et al., Cerebral Cortex, 2019).

By using a novel technique for analysis of synapses in brains of individuals affected by neurodegenerative disorders, we demonstrated that the loss of synapses in Alzheimer's disease is linked to the degradation of synaptic neural adhesion molecules (Leshchyns'ka et al., 2015; featured at the Medical News website and others, the front page of the UNSW website and in the Newsletter (Summer 2015/16) of the Australia and New Zealand Society for Cell and Developmental Biology). Currently, we analyse mechanisms of this loss and develop strategies that can be used to prevent it. We also investigate changes in neural cell adhesion in other neurodegenerative disorders such as Parkinson's disease and motor neuron disease and use new transgenic mice to model abnormal function of neural cell adhesion molecules in these disorders and study its effects on the brain.

REFERENCES

- » Leshchyns'ka I et al. 2015, 'Aβ-dependent reduction of NCAM2- mediated synaptic adhesion contributes to synapse loss in Alzheimer's disease', Nature Communications, 6:8836.
- » Sheng L et al., 2015, 'Neural cell adhesion molecule 2 promotes the formation of filopodia and neurite branching by inducing submembrane increases in Ca2+ levels', *Journal of Neuroscience*, 35:1739-52.
- » Sytnyk V et al, 2017, 'Neural cell adhesion molecules of the Immunoglobulin superfamily regulate synapse formation, maintenance, and function', *Trends in Neuroscience*, 40:295-308.





APPROVED EXTERNAL SUPERVISORS

HDR may also be undertaken with approved external supervisors located in institutions affiliated with the School of BABS. Students can contact external supervisors directly for information on available projects. Please note that it is BABS policy that a BABS academic must be assigned as joint supervisor.

FREQUENTLY ASKED QUESTIONS

1. I'm interested in postgraduate study in BABS - where do I begin?

First and most importantly, it is essential that you identify an appropriate academic supervisor in BABS and obtain their agreement prior to submitting an application for postgraduate study.

It is recommended that you peruse the School's current research clusters to obtain an understanding of the areas of study available within BABS. You should try and align your topic of interest with the research area of one of the affiliated academics. Contact details for individuals can be found in the BABS Academic and Research Leaders Directory. This booklet has also been designed to capture this information within the Research Supervisors sections. We also produce a BABS Honours Information Booklet which lists supervisors and honours projects (these projects may be relevant for prospective MPhil students).

In your email, it is recommended that you:

- a. Identify which research area you are interested in, and why
- b. Indicate which term you intend on commencing (Term 1, 2 or 3)
- c. Advise your availability times for an interview
- d. Attach a copy of your CV and academic transcript
- e. Confirm you have available funding to cover both living expenses and the tuition fees
- f. Indicate clearly that you have appropriate visa status (or will apply for same)

NB: If you submit an application for postgraduate study without a nominated supervisor, it is likely to be declined.

Identifying and negotiating with prospective supervisors is up to you. Your ideal supervisor will be knowledgeable in your topic of interest, have good research skills and experience, and be someone you feel you can work well with. Choosing the right supervisor is very important. It is recommended that you meet and/or correspond with those academics in BABS that you feel have expertise in your area of interest to identify the best person to ask to be your supervisor. Note, however, that as most of our academic staff receive between 50 and 200 requests from prospective postgraduate students each year, please understand they may not always be able to assist you.

2. What are the visa requirements?

Overseas postgraduate students must obtain an appropriate visa that allows them to study in Australia. Please refer to the <u>Australian Dept of Home Affairs</u> and <u>UNSW visa information</u> to ensure you are aware of the specific requirements in regard to obtaining the correct visa. This process can be very time-consuming, so please make sure you know what is required.

FREQUENTLY ASKED QUESTIONS

3. What are the academic entry requirements?

All entry requirements for the particular degree must be met (for instance, an Honours undergraduate degree is generally a pre-requisite for entry into a PhD program). Admission criteria are an academic matter and you will need to provide evidence that you have attained the appropriate pre-requisites for the degree you are applying for. Requirements for each degree is provided in this booklet and can also be found online.

4. What scholarships are available?

Please note that the School does not provide scholarships to support either living expenses or tuition fees for postgraduate students. The cost of living in Sydney can be high, and part-time employment may not be readily available or permitted under some visa conditions. Accordingly, it is your responsibility to ensure you have adequate funding to cover both of these necessities. The University of NSW does provide some postgraduate scholarship opportunities, which are detailed in this booklet and can be found on the UNSW Graduate Research Scholarships website and the UNSW Scholarships website.

BABS has some <u>scholarship opportunities</u> available for students once they are enrolled.

5. What are the English language requirements?

Full information on the English language requirements of this University is available <u>here</u>.

6. How do I apply?

Once you have found a supervisor, please see the <u>Graduate Research Submit an Application webpage</u> if you are applying for PhD, MSc or MPhil. Students can apply for the UNSW scholarships during their admission application. HDR admission and scholarship FAQs are available <u>here</u>. Please note that the Graduate Research School administers postgraduate research study only. Undergraduate programs are administered by The <u>UNSW Nucleus Student Hub</u>.



