

University of Oxford Laboratory Services Showcase

BioEscalator, 18th May 2023

Sponsored by





1. Introductions

Dr Claire Shingler

Dr Emma Packard

Welcome



Incubator for spin-out and start-up biomedical science companies

- Shared and private CL2 labs
- Entrepreneurial and business support



Today's event



- 24 Flash presentations (2 mins)
- Networking lunch/Q&A
- Housekeeping
- Sponsors: Oxford University Innovation and The Oxford Trust

Oxford University Innovation Limited

Creating global impact from academic discovery and expertise



 Enabling impact from discovery through Licensing, IP and Patents, Spinout formation, Material Sales, Clinical Outcomes, Startup Incubator, Social Enterprise creation, Oxford University Hospitals NHS Foundation Trust

Patent

Licensing & Ventures

• Supporting researchers and external partners to utilise academic expertise and technical services

Consulting Services Enabling ventures to grow from concept to maturity through Seed Funds, Oxford Investment Opportunity Network, Spinout Equity Management

Funding, Investment & Management

Consulting Services

Enabling Oxford expertise to have impact in the wider world





Providing external organisations with access to University expertise & facilities.



Helping academics identify and manage consulting opportunities.



Supporting Departments in arranging external services (including consultancy) work. **2,356 academic consultants** registered with OUI Consulting Services

- providing advice and expertise to organisations worldwide.



OXFORD UNIVERSITY

Contact us

in

emma.packard@innovation.ox.ac.uk

sally.sheard@innovation.ox.ac.uk

enquiries@innovation.ox.ac.uk

www.innovation.ox.ac.uk

linkedin.com/company/oxford-university-innovation

twitter.com/OxUInnovation







2. Advanced Proteomics Facility

Dr Marjorie Fournier



Advanced Proteomics Facility

Dr Marjorie Fournier, Manager

OU Facilities Showcase Bioescalator Oxford, 18th May 2023





APF Proteomics Applications



APF Proteomics Applications



"Key methodologies to identify molecular mechanisms regulating biological processes"





3. OHS Specialised Analytical Service Provider

Dr Rod Chalk









Rod Chalk PhD

Everything you need to know about your expressed protein





Over 20 years experience in the structural characterization of glycoprotein assemblies – from biotherapeutic antibodies to viruses and vaccines



Glycomics



Uncovering the molecular detail of glycan structures: topology, branching & monosaccharide composition



Resolving positional information of glycan attachment to the peptide backbone



structural information

Structural and functional analysis of expressed proteins using top-down, bottom-up & native mass spectrometry



Complexes



Stoichiometry determination for homo and heterocomplexes

Membrane proteins



ADCs



Confirmation of chemical conjugation reactions and quantitation of drug payloads



Resolving conformational changes in native structures



Drug targets

Compound and fragment screening





General enquiries: rod.chalk@cmd.ox.ac.uk Service contracts: sally.sheard@innovation.ox.ac.uk



Find me at the network session

Primary structure Post-translational modification Chemical modification Purity Conformation Functional activity Protein-protein interaction Protein-ligand interaction



4. PX (Protein Crystallography)

Dr Lizbé Koekemoer







PX (Protein crystallography) SRF

Lizbé Koekemoer

18 May 2023 •

Our facility

- Walk-in facility for academic collaborators:
 - Access to crystallography equipment and facilities, including an extensive screening library
 - Diamond sample registration, and tracking of crystals using our Scarab database
 - Crystal storage and shipping to Diamond, and subsequent disposal
 - Tracking of crystals using Scarab database
 - Diamond BAG reporting
- Crystallography as a service for academic and industrial users:
 - Crystallization of novel targets, including trouble-shooting of difficult targets
 - Reproducing published structures and internal projects
 - Hit optimisation and elaboration
 - Protein-Ligand and Protein-Protein complexes
 - LCP crystallography of membrane proteins
 - Structure solution and PDB deposition preparation
 - Project management and reporting
- XChem:
 - Getting crystals XChem-ready Performing XChem screens
 - Data analysis and reporting
- Scientific support and method development, including protein engineering (in collaboration with the Protein Production SRF).
- All data capturing and reporting done using Scarab (Research informatics) and RockMaker web.













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5. Protein Production

Dr. Ellie Williams





Protein Production Small Research Facility

• Dr. Ellie Williams – Protein production facility coordinator

Protein Production Facility - CMD

Construct Design and Cloning

- Help with construct design available
- High throughput 96 well plate cloning
- Access to 150+ vectors for *E. coli* / Insect / Mammalian systems

Of Note

- Continuous pipeline into RI, MS, Biophysics and Crystallography SRF units
- Keen to work with academia and industry

Would like to meet...

- Researchers looking for both short and longer term collaborations
- Anyone looking to talk over ways we could help them across the CMD SRF pipeline



Protein Expression

- *E. coli,* Insect and Mammalian expression platforms in use
- 80+ *E. coli* cell strains available
- Virus generation for baculoviral strategies

Protein Purification

- Expertise in protein purification for multiple final applications
- Development of purification strategies
- Experience in soluble, refolding, secreted and endotoxin free purifications



Contact: Eleanor.Williams@cmd.ox.ac.uk



Biophysics and Biochemistry Research Group



Full range of biophysical instrumentation for hit identification and validation





SPR









ITC

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BLI



Protein stability & aggregation



RapidFire MS

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Contact: oleg.fedorov@cmd.ox.ac.uk



6. NDM Centre for Translational Proteomics

Dr Roman Fischer

NDM Centre for Translational Proteomics







DisPro (Discovery Proteomics)







- 🌞 Thermo QExactive
- Thermo Orbitrap Fusion Lumos
- Thermo Orbitrap Ascend + FAIMS
- Bruker TimsTOF PRO + Evosep One

 \Rightarrow Staff: 6 \rightarrow 8

♦ Assets: £8 → 10 M

CellPro (Cellular Proteomics)



- Bruker TimsTOF FLEX ESI/MALDI
- Bruker TimsTOF SCP + Evosep One
- Waters Cyclic IMS HDX (coming soon)



Clinical

Proteomics

roman.fischer@ndm.ox.ac.uk

Laser Capture Microscope CellenOne (single cell dispenser) Bravo liquid handler Covaris

- \bullet SRF \rightarrow cost recovery
- 🌻 # publications: 20-30 / year



~11/2023



7. Nuclear Magnetic Resonance (NMR) Spectroscopy

Professor Christina Redfield



BioEscalator Facilities Showcase

Nuclear Magnetic Resonance (NMR) Spectroscopy Facility in the Department of Biochemistry

18 May 2023



Solution NMR Facility in Biochemistry

- The facility is open to all members of the Biochemistry Department, Medical Sciences Division and the wider University.
- Access to the NMR facility for external companies (i.e. Biotech, Pharma) is also very welcome and this can be arranged via Oxford University Innovation.
- The NMR facility is located in the South Parks Road Science area and has NMR spectrometers operating at ¹H frequencies of 500, 600, 750 and 950 MHz.
- All spectrometers are equipped with Bruker consoles and high-sensitivity inverse ¹H/¹³C/¹⁵N triple resonance 'cryo'-probes. The 600 MHz is also suitable for ¹⁹F NMR. The 950 has a fully-automated sample changer that can handle racks of 96 NMR tubes.

Solution NMR Facility in Biochemistry

- Training/advice is available for anyone interested in using NMR either as independent users or through collaborations with Redfield/Schnell research groups. A 'service' could also be available.
- Remote operation of the NMR spectrometers is possible. We are part of the Horizon 2020 Remote-NMR project working on developing a common protocol.
- Experiments can be set up with help of facility staff via an interactive Teams/Zoom session or remote user can operate the spectrometer without assistance from facility staff. Samples would need to be shipped/delivered to the Biochemistry Department.

The 950 MHz NMR spectrometer is equipped with a SampleJet sample changer which enables higher throughput experiments.





These might include ligand screening for drug discovery via either ligand or protein detect methods.

The SampleJet can accommodate 3mm or 5mm 'standard' NMR tubes in the ~95 positions around the carousel or racks that hold up to 96 3mm or 5mm short NMR tubes. The samples in the racks can be temperature controlled prior to and after data acquisition. Automation of data collection is done using IconNMR.





Solution NMR Facility in Biochemistry

- The Redfield and Schnell research groups have expertise in biomolecular NMR, particularly studies of structure, dynamics and interactions of proteins in solution.
- The NMR spectrometers are also suitable for characterization of small organic compounds (TOCSY, HSQC, HMBC etc), kinetic measurements (substrate turnover in enzymatic reactions), ligand binding studies (ligand-observe STD NMR, protein-observe titrations), and metabolomics.

Contact information

 Please contact: <u>christina.redfield@bioch.ox.ac.uk</u> if you'd like more information about our NMR facility or to discuss whether NMR might be of use in your research & development projects.



8. COSMIC cryo-Electron Microscopy Facility

Dr Rishi Matadeen
CENTRAL OXFORD STRUCTURAL MOLECULAR IMAGING CENTRE

Structural Biology Solutions



3.88 Å structure of Cytoplasmic Polyhedrosis virus by cryoelectron microscopy

Courtesy of Xuekui Yu, Lei Jin & Z. Hong Zhou, University of California, Los Angeles, USA Cellular Biology Solutions Discover life's cellular architecture in 3D



Volume rendering of the threedimensional architecture of a dividing yeast cell

Courtesy of Sriram Subramaniam, National Institutes of Health, Bethesda, USA CENTRAL OXFORD STRUCTURAL MOLECULAR IMAGING CENTRE



Equipment at Cosmic

• 3 Thermo Scientific Mark 4 Vitrobots for rapid freezing of samples

- Dedicated humidity-controlled preparation lab
- Several TEMs and a FIB SEM

CENTRAL OXFORD STRUCTURAL MOLECULAR IMAGING CENTRE



Central Oxford Structural Molecular Imaging Centre

Highlights

• Streamlined Single Particle Analysis workflow:

-ß-galactosidase resolved to 1.95Å, 4000 movies, 6 hour data set collection

• Fully Supported Sample Preparation and Data Collection

- Established Industrial customer pipeline

• Comprehensive Training from Thermo Fisher Scientific and COSMIC Support staff

Central Oxford Structural Molecular Imaging Centre

COSMIC TEAM CONTACTS

Prof Matthew Higgins, COSMIC Scientific Director <u>matthew.higgins@bioch.ox.ac.uk</u>

Dr Rishi Matadeen, COSMIC Microscope Facility Manager <u>rishi.matadeen@path.ox.ac.uk</u>

Emma Packard, OUI Project Manager emma.packard@innovation.ox.ac.uk

COSMIC cryo-EM facility website.



9. Oxford Particle Imaging Centre (OPIC)

Professor Jon Grimes



- OPIC offers a wealth of high resolution imaging resources for structural analysis by electron microscopy
- Single Particle analysis cryo-EM and cryo-electron tomography (cryo-ET) in CL3 containment laboratories
- FIB-SEM microscope for the milling of thin cellular lamella for in-situ cryo-ET in CL3 containment laboratories
- Unique in Europe (and perhaps elsewhere) in having cryo-EM in contained CL3 laboratories.

Aquilos2 FIB-SEM



With integrated fluorescence light microscopy for correlative imaging. Within HG3 suite.

<u>Collaborative</u>

projects...from **crystals** to **cell** samples; from picornato coronaviruses.

To mill thin 120nm Lamella from frozen cells



Glacios TEM

• AFIS.

- Latest EPU, Tomo5.
- HG1-2 pathogens..

Screening of grids to optimise sample preparation

Titan Krios G3i TEM



- Falcon IVi/Selectris X.
- 550-750 movies an hour
- AFIS, FFI.
- EPU, SerialEM, cryo, Tomo5, PACE-tomo.
- HG1-3.

High resolution SPA cryo-EM and cryo—ET

Blot /plungefreezing devices



- GP2, Leica (HG3).
- 2 Vitrobots.
- 3 manual plunging devices, some at HG3.
- CP3, Gatan (HG3).

Freezing of grids for SPA cryo-EM and cryo—ET

To conclude

- OPIC facility offers a competitive range of equipment for electron microscopic structural analysis, including a Titan Krios with Falcon-4/SelectrisX detector within a hazard group
 3 level (HG3) facility unique in Europe.
- Oxford, UK; website: <u>opic.ox.ac.uk</u>
- Access is open for both academia and industry
- Academic users from EU countries have the option to apply for **Instruct funding**
- Extensive collection of equipment and very enthusiastic and research focussed staff
- Email <u>emsupport@strubi.ox.ac.uk</u> with any enquiries!



10. Dunn School Electron Microscopy Facility

Dr Errin Johnson

Dunn School Electron Microscopy Facility

BioEscalator University Services event

Dr Errin Johnson



Dunn School Electron Microscopy (EM) Facility

- Based in the Dunn School of Pathology at Oxford University, the EM Facility has everything you need for any type of biological EM. For example, we offer:
 - Quick turnaround screening & QC checks of purified samples, including extracellular vesicles (EVs), Virus-like Particles (VLPs), proteins and α-synuclein & Amyloid Beta
 - High resolution imaging of cells, tissues and whole organisms to assess morphological and ultrastructural changes due to disease, genetic manipulation or drug treatments
- Advanced EM techniques are also supported, including volume EM, immunogold labelling, elemental mapping and correlative light and electron microscopy (CLEM)
- We provide expert advice on which EM techniques are right for your project, as well as support in image interpretation and troubleshooting
- The EM facility is open to both research and industry users
- We offer training, as well as full service options
- For more details, please visit our website: www.dunnschoolbioimaging.co.uk
- Please get in touch with me to discuss how EM can benefit your work: <u>errin.johnson@path.ox.ac.uk</u>









11. Dunn School Light Microscopy Facility

Dr Alan Wainman

dunn school



Instruments

Fluorescence

'Super-res' confocal



Evos M7000 (long time-lapse, Plate imager, histology)



Zeiss 880 Airyscan (Airyscan Fast- FCS)

Confocal



Olympus FV1000 (With B&H FLIM: Pulsed 488nm laser)



Zeiss fluorescence



Olympus SoRa spinning disc (405nm photomanipulation unit)



Olympus FV1200 (SIM scanner..DNA damage)







Ease of access

Many commercial users have already used the Dunn School facility. We have experience with many samples and projects.



EM and LM

Both LM and EM work close together and we have a proven track record of successful commercial projects requiring



Parking-

Parking available at Dunn School

www.dunschoolbioimaging.co.uk



alan.wainman@path.ox.ac.uk





12. Cellular Imaging Core Facility

Dr James Bancroft

Cellular imaging Wellcome Centre for Human Genetics

Cellular Imaging Core Facility

James Bancroft, PhD



Who & where?

- 1 minute away from the Innovation Building
- Team of 2 (soon to be 3)
- Over 20 years of combined microscopy experience
- Expertise in sample preparation and imaging and analysis from from tissues and cells to single molecule imaging











James Bancroft

Ed Drydale

How do we work?

- We offer our services at all project stages, from experimental design to sample preparation, through to image acquisition and analysis
- We generate custom analysis pipelines leveraging the latest AI segmentation tools
- We can image samples for or with users or users can be trained to access systems independently
- We help users achieve the right kind of resolution



Our systems

Olympus/Evident 2 cam SoRa (super-res spinning disc) with ScanR high content package (image based cytometery)

Zeiss LSM900 with Airyscan2 (super-res point-scanning confocal) + direct processing workstation

Zeiss spinning disc confocal (3 channel)

Zeiss Laser Capture Microdissection (LCM) with fluorescence and colour

Leica SP8 confocal with white light laser

Leica SP8 FALCON - lifetime imaging

Leica widefield with fast filters and colour imaging (x2)

Aperio colourometric slide scanner (5 slides at a time)

Coming soon - Olympus/Evident VS200 - 200 slide capacity with fluorescence, colour and polarisation imaging

Elyra PS.1 dSTORM/SMLM + TIRF - dual camera

Elyra 7 lattice SIM with 3D dSTORM/SMLM and TIRF - dual camera (Joint acquisition with Zeiss CoE) EVOS M5000 - bench top fluorescence and colorimetric imaging

3 X Titan ultra-high spec image analysis workstations with Arivis Vision4D (x2 licences), Imaris, Zen, LasX, ScanR workstation for offline analysis of high content imaging

Example projects

Live cell uptake assays

High content imaging & Image based cytometry



Cell tracking & wound healing



Laser Capture Microdissection

Organoid imaging

Fluorescence Lifetime Imaging (FLIM)/Biosensors









Contact cellular-imaging@well.ox.ac.uk







13. Micron Imaging Facility

Dr Deidre Kavanagh

Micron OXFORD

Home

About us >

Our Technology >

Research >

Access >

Education and Outreach >

News and Events





Dorothy Crowfoot Hodgkin Building

MICRON BIOIMAGING FACILITY

Access to advanced imaging technology Including bespoke development systems With expert guidance and support for your project

BOOK A MICROSCOPE

micron@bioch.ox.ac.uk www.micronoxford.com @MicronOxford





The Micron Team



Prof. Lothar Schermelleh Micron Director & Academic Lead



Dr Deirdre Kavanagh Micron Manager



Dr Niloufer Irani Micron Ass. Manager





Micron Technology



• 8 turn-key instruments and 2 development instruments

Access to

Confocal microscopy Widefield microscopy High Content imaging Single Molecule Tracking and Localization Structured Illumination Microscopy AI Analysis Live and fixed imaging Secure data storage









Micron Technology





Micron Technology Focus



Olympus ScanR High Content Screening Widefield Microscope



Applications

- Quantitative Image-based Cytometry (QIBC)
- Cell cycle assays
- Protein localization and colocalization
- Gene expression
- Intracellular transport
- Infection assays
- Cell migration assays
- Rare event analysis


















Micron Courses



- Annual microscopy workshops and courses including sample preparation and data analysis workshops.
- Bespoke microscopy courses available on request.
- In person and remote courses available.
- Host seminars and taster sessions.
- Late 2022 4 day super-resolution course (SIM).



Micron Access

- Access managed through Stratocore PPMS core facility management software.
- Not only user bookings but instrument logging and tracking to ensure minimum down time.
- SRF FastTrack visitor process to the Department of Biochemistry.
- We provide as much training and support as required for your project.
- Independent route following training on the instruments.
- Service route where Micron perform data acquisition and processing.

Access Rates:

- Flat rate per hour or per day depending on your access requirements.
- Options to include data storage (micron server including tape back-up) and consumables.





Thank you for listening



Micron Manager: Deirdre (Dee) Kavanagh Micron Assistant Manager: Niloufer Irani Academic Lead: Lothar Schermelleh

Website: Twitter: Email: www.micronoxford.com @micronoxford micron@bioch.ox.ac.uk







14. Oxford-ZEISS Centre of Excellence

Dr Helena Coker

Microscopy in IDRM, KIR, and Oxford-ZEISS CoE









Helena Coker, PhD Microscopy Facility Manager

Expert Skills: Light Sheet / Lattice Ligh Sheet SIM Kseniya Korobchevskaya, PhD Microscopy Facility Manager

> Expert Skills: Confocal, Super-resolution Airyscan and SIM



Jacky Ka Long Ko, PhD Advanced Image Analyst

Expert Skills: 3D cell analysis, Volume tracking Machine learning

Email us:

imaging-facility@kennedy.ox.ac.uk

www.kennedy.ox.ac.uk/platforms-and-technologies/advanced-microscopy-centre

Microscopy in IDRM, KIR, and Oxford-ZEISS CoE





Light Sheet ZEISS LLS7, Z1 , LaVision UMII

- Low phototoxicity
- (sub)cellular fast live acquisition (LLS7)
- Multiview live or aqueous cleared (Z1)
- Large organically cleared tissues (UMII)

Confocal ZEISS 980 w/ Airyscan 2

- High spatial resolution •
- Live (CO2 and temp control)
- FLIM, FRAP and FCS





Slide Scanner **ZEISS Axioscan 7**

- Brightfield and Fluorescence
- 7 LEDs •
- Up to 100 slide automated scanning

Widefield Leica w/ Phenocycler (CODEX)

- Easy to use
- Spatial phenotyping up to 100 channels)
- Live (temperature control)



K. Korobchevskaya, A. Magnussen











15. Microscopy in the Department of Oncology

Dr Rhod Wilson



Microscopy in the Department of Oncology

Rhod Wilson, Scientific Technologies Manager





What we offer

AVAILABLE FACILITIES

- Andor Dragonfly
- Zeiss LSM 710
- Zeiss LSM 780
- Zeiss LSM 880
- Nexcelom Celigo
- Nikon Ni-E
- Nikon Ti-E
- Nikon 90i
- Nikon TE2000-E
- Imaris Workstation

AVAILABLE TECHNIQUES

- Airyscan Super-Resolution Microscopy
- Laser Scanning Confocal Microscopy
- Spinning Disk Confocal Microscopy
- Multi-photon Excitation Microscopy
- Widefield Microscopy including fluorescence, DIC and Phasecontrast
- Plate Scanning Image Cytometer
- Image Analysis Expertise

Celigo Imaging Cytometer





Contact us

Rhodri Wilson – SRF Manager

Rhodri.wilson@oncology.ox.ac.uk 01865 617421 ORCRB, Roosevelt Drive OX3 7DQ

Visit our website:



www.oncology.ox.ac.uk/research/srf/microscopy





16. The Don Mason Facility of Flow Cytometry

Dr Robert Headley

The Don Mason Facility of Flow Cytometry



DR ROBERT HEDLEY (DPHIL)

Flow Cytometry Facility Manager at the SWDSOP robert.hedley@path.ox.ac.uk



VASILIKI TSIOLIGKA (MSC)

Flow Cytometry Specialist at the SWDSOP Vasiliki.tsioligka@path.ox.ac.uk



Cell Sorting Service



Cell analysis



Consultations (Cell prep, panel design, and staining protocols) - Free



Analyser Training – Free



FlowJo, Cytobank, or IDEAS training -Free

Our Location



Sir William Dunn School of Pathology, Level 1



Flow cytometry allows the analysis and/or sorting of single cells or small particles in a liquid suspension, based on their refraction of light or fluorescence.





Our Extended Support



Jump to:

Start the X20 Setting up an experiment (tubes) Setting up an experiment (HTS) Daily cleaning procedure Shut down or standby Troubleshooting

Start the X20

To start X20 switch on the green button on the right hand side of the X20. If it is already on, make sure that it is on 'Standby' and 'Low'.



Start the computer

- User: Operator
- Password: facs

Please visit our website for detailed information and booking





Cytoflex LX

lur mast accessible machine for new flow cytome with APD detectors that specialise in measuring th within the for red scale and offers superior res	triate, the Cytoffex IX is fitted acressent emission spectra sution to PMF detectors.	
optios	~	
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sage	~	

Use and		
applications		

+ What cells can I run on this machine?	
---	--

+ How long does data acquisition take?

+ How much does data acquisition cost?



https://www.oxforddmflowcytometry.com/



17. Flow Cytometry Facility, Experimental Medicine

Dr Helen Ferry

Experimental Medicine Division, NDM (Translational Gastroenterology Unit)

Flow Cytometry Facility

John Radcliffe Hospital

Helen Ferry (helen.ferry@ndm.ox.ac.uk)







Where are we?



Image credit: OHS NHS Foundation Trust

Experimental Medicine Division: TGU/RMU



Other departments/customers



Other departments/customers



Facility equipment – conventional analysers

BD LSRII SORP cell analyser - 18 fluorescent parameters

- 405nm violet laser (100mW) 8 fluorescent parameters
- 488nm blue laser (80mW) 2 fluorescent parameters
- 532nm blue-green laser (150mW) 5 fluorescent parameters
- 642nm red laser (40mW) 3 fluorescent parameters



• High Throughput Sampler available for sample acquisition from 96- and 384-well plates

BD LSRFortessa cell analyser - 14 fluorescent parameters

- 405nm violet laser (50mW) 6 fluorescent parameters
- 488nm blue laser (50mW) 5 fluorescent parameters
- 640nm red laser (40mW) 3 fluorescent parameters



Facility equipment – cell sorter

BD FACSArialII cell sorter - 17 fluorescent parameters

- 405nm violet laser (50mW) 7 fluorescent parameters
- 488nm blue laser (20mW) 2 fluorescent parameters
- 561nm yellow-green laser (100mW) 5 fluorescent parameters
- 633nm red laser (18mW) 3 fluorescent parameters
- Sorting into tubes (0.2, 0.5, 1.5 and 15ml), plates (TC, PCR, Terasaki), microscope slides etc.
- Housed in a Class I MSC
- Operated by facility staff only



Facility equipment – spectral analyser

Cytek Aurora spectral cell analyser

- 355nm UV laser (20mW)
- 405nm violet laser (100mW)
- 488nm blue laser (50mW)
- 561nm yellow-green laser (50mW)
- 640nm red laser (80mW)



Advantages of Aurora spectral flow cytometry:

- High-parameter panels (30+ colours) using standard commercially available dyes
- Autofluorescence (AF) subtraction to improve resolution for cell types with inherently high cellular AF or for applications such as intracellular cytokine staining
- Violet side-scatter detection for improved small particle detection
- Exclusion of red blood cell contamination through differential SSC gating
- Superior detection of red, far-red and infra-red emitting dyes

OMIP-069: Forty-Color Full Spectrum Flow Cytometry Panel for Deep Immunophenotyping of Major Cell Subsets in Human Peripheral Blood. LM, Lannigan J, Jaimes MC.Cytometry A. 2020 Oct;97(10):1044-1051

What can we offer you?

- High quality sorting service from operator with > 15 years experience sorting a range of tissue and cell types (PBMCs, bone marrow, spleen, colon, lymph node FNA, cerebral spinal fluid, liver FNA, bronchial alveolar lavage, lung, sputum, endometrium, spider caccoons, sputum, breast milk, various cell lines etc.)
- Training for new users including theory and practical
- Help and advice with panel design (more than 4 years experience designing panels specifically for spectral flow cytometry)
- Help troubleshooting experiments
- Acquisition of samples on the Aurora

Contact details

Facility Manager: Dr Helen Ferry (she/her)

helen.ferry@ndm.ox.ac.uk

http://www.expmedndm.ox.ac.uk/flow-cytometry-facility

TGU/Experimental Medicine Division NDM Level 5, Room 5605 John Radcliffe Hospital Headington Oxford OX3 9DU



If you would like to know more about the facility, I am very happy to discuss your requirements and answer your questions, so please drop me an email

OUI contact: <u>Sally.Sheard@innovation.ox.ac.uk</u>



18. The WIMM Flow Cytometry Facility

Dr Paul Sopp





The WIMM Flow Cytometry Facility

MRC Weatherall Institute of Molecular Medicine, John Radcliffe Hospital, Headington, Oxford, OX3 9DS





John Radcliffe Hospital site map



The Team

Paul Sopp Kevin Clark Sally- Ann Clark Craig Waugh

Flow Cytometry Facility







Cell Analysers

Analyser	Lasers	Parameters
3 x Attune NxT	4	14
Fortessa	4	15
Fortessa X20	4	16
Fortessa X20	5	18
Symphony A5	5	30

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Cell Sorters

Analyser	Lasers	Parameters
Aria III	4	15
2 x Fusion	4	16
Symphony S6	5	30
Sony MA900	4	12




What we can offer

- Cell sorting into tubes and microplates including single cell sorting for cloning
- Self-assisted and assisted cell analysis
- Training theory and practical
- Panel design advice and testing
- Large scale cell sorting using 2 or more sorters for very large samples
- Collaborator access into the Facility (24h access)
- Projects including use of other WIMM Facilities including the Single Cell Facility services



• Formal service agreements facilitated by OUI (sally.sheard@innovation.ox.ac.uk)





19. Flow Cytometry Facility IDRM

Dr Jesus Reine



FLOW CYTOMETRY FACILITY IDRM Old Road Campus, Roosevelt Drive, Oxford OX3 7TY Contact: <u>Jesus.Reine@paediatrics.ox.ac.uk</u>



FLOW CYTOMETRY FACILITY IDRM – Who we are



Superusers Committee – Team highly trained in flow cytometry



Superusers competences

- Flow cytometry (conventional and/or spectral)
- Laser safety training
- Laboratory safety induction
- Collaborations and expertise
- Access to our facilities and instruments



We have 11 instruments with 4 available to the public.

Details about our service and instruments can be found in the QR brochure.



Acquisition services with Sony spectral cytometer ID7000

Instrument	Location	Lasers	Ex wavelength (nm)	Detectors	Sample loading options
ID7000	Ground	5	355/405/488/561/637	147	24 tubes rack, 96 and
	floor				384 well plate
ID7000	Level 1	4	405/488/561/637	112	24 tubes rack, 96 and
					384 well plate
ID7000	Level 2	5	355/405/488/561/637	147	24 tubes rack, 96 and
					384 well plate









FLOW CYTOMETRY FACILITY IDRM – Where we are





FLOW CYTOMETRY FACILITY IDRM Old Road Campus, Roosevelt Drive, Oxford OX3 7TY Contact: <u>Jesus.Reine@paediatrics.ox.ac.uk</u>





20. Genome Engineering Oxford (GEO)

Dr Joey Riepsaame

An introduction to the Genome Engineering Oxford (GEO) Facility



Joey Riepsaame

O1865 285489 joey.riepsaame@path.ox.ac.uk Dunn School of Pathology – OMPI 3





Next-Gen DNA editing: Prime Editing



- Editing without introducing double-stranded DNA breaks!
- Suitable for knock-outs, knock-ins and SNPS
- DNA repair template is incorporated into the prime editing guide RNA (pegRNA)
- Specificity: +++

Conventional DNA editing: Cas9 vs Cas12a



- Recognizes NGG PAM
- 2-component guide (crRNA + tracrRNA)
- Multiplexed guides? NO
- Specificity: +
- Generates blunt-ended cuts



- Recognizes TTTV PAM
- 1-component guide (crRNA only)
- Multiplexed guides? YES
- Specificity: ++
- Generates staggered ("sticky") overhangs

NB: DNA editing occurs through introducing double-stranded DNA breaks

Genome-Wide CRISPR Library Screens



Genome-Wide CRISPR Library Screens: Humagne Platform

а



DeWeirdt et al. - Nat Biotechnol. (2021)

GEO Services provided

Current services:

Advice on gene editing strategies	Free
Custom-made clonal cell lines: knock-outs	Starting at £3000 per line
Custom-made clonal cell lines: point mutations, knock-ins	Starting at £4000 per line
Pooled gene edited cells custom knock-outs, SNPs/knock-ins*	Starting at £750 per line

Upcoming services (summer 2023):

Multiplexed genome-wide CRISPR library screening**	Starting at £10K / sample
CRISPR library prepping and NGS validation	Starting at £2K / library

*Price does not include the costs of HDR template

** Platform: Multiplex Humagne C and D libraries (John Doench, Broad Institute)





See you soon at the GEO Facility!

Location: Dunn School of Pathology – OMPI 3



Joey Riepsaame (PhD) Head of GEO joey.riepsaame@path.ox.ac.uk



Georgie Fisher (MSc) Cell line engineering georgie.fisher@wolfson.ox.ac.uk



Richard Wallbank (PhD) CRISPR library screening richard.wallbank@path.ox.ac.uk

For more info, please visit:

http://www.path.ox.ac.uk/content/genome-engineering-oxford-geo

2 1865 285489





Next-Gen DNA editing: Prime Editing



- Editing without introducing double-stranded DNA breaks!
- Suitable for knock-outs, knock-ins and SNPS
- DNA repair template is incorporated into the prime editing guide RNA (pegRNA)
- Engineered pegRNAs and PEmax systems now available at GEO



21. Cellular High Throughput Screening

Dr Daniel Ebner



nature genetics

A genome-wide CRISPR screen identifies CALCOCO2 as a regulator of beta cell function influencing type 2 diabetes risk

Antie K. Rottner @¹. Yingving Ye @^{2,9}. Elena Navarro-Gu

https://www.cmd.ox.ac.uk/ research/high-throughputscreening

"Classic" microtiter plate-based, phenotypic screening using arrayed

- 1) small molecule (Anti-cancer, FDA Approved, Active and 50K Diverse Set Libraries)
- 2) CRISRP libraries (CRISPRn, CRISPRi and CRISPRa)
- 3) siRNA libraries (Yes, siRNA is still a thing)



sequenced for synergy"

Ebner Lab research focusses on identifying novel combinatorial targets in GBM, specifically, Glioblastoma Stem Cells



Genome-wide CRISPR/Cas9 loss & gain of function screening (CRISPRn, CRISPRi and CRISPRa)

XCELLOMICS

Translating Novel Biology Into Drug Discovery

Collaboration between Exscientia/CMD – TDI to accelerate early-stage drug discovery research in academia and then partner with industry

https://www.xcellomics.com/



About the lowerships Dublish with us

hwen Tay, Justin P. Whalley, Sally A. Cowley, Ben Davies, Julian C. Knight 🗠

are > scientific reports > articles > artic

Phenotypic Screening; Automated Microscopes

IN Cell Analyzer 6000 (Cytiva) Line scanning confocal



405 UV 488 BLUE 561 GREEN 642 RED BRIGHTFIELD Simulated Phase & DIC Temp control/CO2/Humidity Time series; many modes, fast to slow Liquid Handling (add drug then image) HWAF/SWAF 2X, 4X, 10X, 20X075, 40X075, 60X095 DEVELOPER TOOLBOX image analysis

Opera Phenix (Perkin Elmer) Spinning Disc Confocal



405 UV 488 BLUE 561 GREEN 640 RED BRIGHTFIELD Simulated Phase Temp control/CO2 Time series HWAF 1.25X, 5X, 10X, 20X, 40X, 63X Including water immersion objectives 4 Cameras for speed

HARMONY image analysis software

Complimentary Instruments & Image Analysis Software

Enabling a wide range of applications; 2D, 3D, fixed, live, plates, slides, dishes, basic or custom image analysis pipelines;



<u>Throughput</u>

Many plates can be loaded automatically using robotic arms Experience

>20yrs experience of High content/throughput imaging & analysis applications

Functional Genomics



Chemogenomics and Drug Screening



Any assay that can be miniaturised to 96, 384, or 1536well plates can be automated on one of our three liquidhandling workstations.

Compound libraries we hold include chemogenomic sets, chemical probes, FDA- and EU-approved drugs, and compound fragments.







22. Dunn School Surface Plasmon Resonance (SPR) Facility

Dr Mikhail Kutuzov

Dunn School Surface Plasmon Resonance (SPR) Facility

Located on the 3rd floor of the OMPI building, Dunn School (South Parks Road)

Biacore T200: one of the most widely used SPR instruments. Provides highly reproducible equilibrium and kinetic binding data with good temporal resolution.



Only very small amounts (few micrograms) of protein are needed to be immobilized on a chip.

Accessible to academic and commercial users

Contact: spr@path.ox.ac.uk

- > Canonical analysis of 1:1 molecular interactions
- Experience in development and application of mathematical models for the analyses of specific cases:
- Enzymatic catalysis (dephosphorylation by SH2-domaincontaining phosphatases)

Goyette et al. (2017) *Sci. Adv.* 2017;3:e1601692 Clemens et al. (2021) *Biophys. J.* 120, 2054–2066

- Ultra-low-affinity interactions (TCR-pMHC) Pettmann et al. (2021) *eLife* 10:e67092
- Protein polymerisation (RAD51 recombinase on DNA) Paoletti (2020) *EMBO J.* 39:e103002
- Bivalent interactions (e.g., antibodies interacting with immobilised antigens)











23. Molecular Biophysics Suite

Dr David Staunton

MOLECULAR BIOPHYSICS SUITE DEPARTMENT OF BIOCHEMISTRY UNIVERSITY OF OXFORD



BioEscalator Facilities Showcase

David Staunton 18 May 2023

Molecular Biophysics Suite

- Organised by Dept and STRUBI and funded by Fell Fund in 2008
- Brought together equipment from various sites to one location in New Biochemistry
- Set up booking system and charging
- Maintenance and servicing
- Training and workshops



Instrument	Technique	Abbreviation
Wyatt HELEOS	Size exclusion chromatography Multiple Angle Light Scatter	SEC MALS
Beckman XL-I	Analytical ultracentrifugation	AUC
Viscotek 802	Dynamic light scatter	DLS
Malvern Vcap	Differential scanning calorimetry	DSC
JASCO 815	Circular Dichroism	CD
Stratagene MC3005PCR	Differential scanning fluorimetry	Thermofluor
Monolith 11.5	Microscale Thermophoresis	MST
Biacore T200	Surface plasmon resonance	SPR
Malvern PEAQ ITC	Isothermal titration calorimetry	ITC
Horiba FluoroMax 4	Fluorimetry	Fluorimetry
Applied Photophysics SX	Stopped flow	Stopped flow
OctetRed 384	Biolayer interferometry	BLI
Agilent Q-TOF	Electrospray ionisation mass spectrometry	ESI-MS



Contact David Staunton

david.staunton@bioch.ox.ac.uk





Dr Mark Hill

Radiation Biophysics: Irradiators Dr Mark A. Hill



23⁸Ptr a-particle irradiator

²³⁸Pu high-LET alpha-particle irradiator (*In vitro*)

Ultrasoft X-ray irradiator (*In vitro*)





320kV @ RRI

X-ray irradiators: in vitro & in vivo













¹³⁷Cs irradiators in vitro

0.662 MeV γ-rays Dose rate:

~0.8 to 10 Gy/min

Used for:

- Radiobiology studies
- Feeder cells
- Hypoxia studies
- (sterilisation)

Mouse environment.

- Anesthetised using isoflurane
- Heated
- Breathing and temperature monitored

Irradiations:

- up to 4 mice irradiated
- Lead used for partial irradiation



320kV X-ray irradiator in vivo

Dose rate: ~1-2 Gy/min

Variable filtration

Radiation Biophysics SARRP: Image-guided preclinical irradiator



CT IP + CT RT guided by CT-MR by CT-MR by CT-MR V CT-MR



Gantry mounted X-ray set Used for.

- Control of cell environment
 - e.g. temp / gassing
- Low dose rate irradiations
- Partial irradiations

320kV @ ORCRB



Development

Stem cell IR of planarian worms



Multi-dose, multi-well IR

Computer controlled lead collimators are remotely moved to vary the number of wells exposed to x-rays so varying the dose across the plate





Multi-well plate

..........

position



²³⁸Pu high-LET α -particle irradiator

Standard conditions:

- 3.3MeV (121 keV/µm) alpha-particles Irradiated cells:
- Dose rate:
 - 20 Gy/min down to <10⁻⁴ Gy/min 4.2 MeV down to < 1 MeV Can vary energy:
 - rticle track





Ultrasoft x-ray irradiator



Standard conditions:

• 1.5 keV X-rays

Can be modified to produced:

- 4.5 keV x-rays
- 0.3 keV x-rays (electron range ~ 4nm)

Partially irradiate cells for DNA damage and repair studies




Dr Mark A. Hill - Radiation Biophysics



More information: Tel: 01865 617056 <complex-block>

Old Read Campus

Email: mark.hill@oncology.ox.ac.uk

Web: https://www.oncology.ox.ac.uk/research/srf/radiation-biophysics



25. Old Road Campus Mechanical Workshop

John Prentice



Old Road Campus Mechanical Workshop

John Prentice and Kyle Hallett Department of Oncology, University of Oxford





About our Facility



- Located in the basement of the Green building (ORCRB), the Department of Oncology mechanical workshop provides support for the research projects within the Old Road Campus, across the other University Departments and beyond.

- We offer a total package from design consultation, computer CAD drawings, material sourcing, manufacture and assembly.
- Our facilities are equipped with state-of-the-art machinery.



What we do





Our modern, fully equipped mechanical workshop enables us to carry out a range of services throughout the course of your project from design to manufacturing. Services include:

- Design services (CAD)
- Manufacture bespoke items
- Repair and maintenance
- Prototyping

- Reverse engineering Measurement services
- Material sourcing
- Technical advice



Work flow



Examples of work:

- Medical sciences
- Imaging microscopes, MRI, etc.
- Radiation

- Cryogenics



• Equipment repair and

- maintenance
- Enclosures and shielding







Projects

Research support

The Process





Contact us



John Prentice John.prentice@oncology.ox.ac.uk 01865 617369 ORCRB, Roosevelt Drive OX3 7DQ

Visit our website:



C DURUC CIP Centre OLD RCAD OLD RC

www.oncology.ox.ac.uk/research/srf/mechanical-workshop



Thank you for you attention.

Sponsored by



Contact details



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- <u>sally.sheard@innovation.ox.ac.uk</u>
- <a>enquiries@innovation.ox.ac.uk

BioEscalator

- <u>www.bioescalator.ox.ac.uk</u>
- <u>https://www.bioescalator.ox.ac.uk/contact/contact-us</u>
- Business Manager claire.shingler@medsci.ox.ac.uk