

## **BBSRC PhD Studentship in Using Directed Evolution to Improve Lentiviral Vector Targeting [Project 2022/07]**

*Radcliffe Department of Medicine (NDCLS), University of Oxford, in collaboration with Oxford Biomedica*

**Application Deadline: Fri 21<sup>st</sup> Jan 2022 (12:00 midday GMT)**

**Project Start Date: October 2022**

### **Supervisors**

Primary Supervisor: Professor Steve Hyde

Secondary Supervisors: Associate Professor James Davies

### **About the Project**

We have developed a novel SIV-based lentiviral vector that directs highly efficient *in vivo* gene transfer to the lungs (PMID: 27852956). The vector incorporates the fusogenic F protein and sialic-acid (SA) binding HN protein from murine parainfluenza (Sendai) virus. Other members of the *Paramyxoviridae* family, of which Sendai virus is a member, have distinct F and HN-like proteins with alternate fusion activities and SA binding properties allowing them to infect alternate respiratory targets. The aim of this project is to generate improved lentiviral vector envelope proteins for enhanced *in vivo* gene transfer via the directed evolution of F and HN sequences.

The project involves the generation of error-prone and/or DNA-shuffled libraries of *Paramyxoviridae* envelope protein genes, the creation of viral vectors that encode their own envelope sequences and the screening/selection of enhanced lentiviral pseudotypes. The project will involve extensive viral vector molecular biology, vector production, and *ex vivo* and *in vivo* gene transfer. The evolution of the libraries will be monitored by conventional and next generation DNA sequencing. Overall, this project is an opportunity to both understand and manipulate the biological interaction of lentiviral vector envelopes with cell surface receptors resulting in improved viral vector targeting and tropism.

### **About the BBSRC Collaborative Training Partnership in Advanced Bioscience of Viral Products (ABViP)**

This PhD studentship is part of the Biotechnology and Biological Sciences Research Council (BBSRC) Collaborative Training Partnership (CTP) in Advanced Bioscience of Viral Products (ABViP). The [ABViP CTP](#) is a comprehensive, multidisciplinary training programme designed to deliver the next generation of bioscience leaders who will advance research on the underpinning bioscience of viral products for future gene therapies and vaccines. Led by Oxford Biomedica and involving both UCL and University of Oxford, CTP students will have access to a wide ranging portfolio of training opportunities at the Partner sites including taught courses and case studies designed to complement the doctoral research. Students trained through the ABViP CTP will gain a holistic insight into the research and development activities required to develop the medicines of the future, with the ability to see the world of medicines development through both an academic and industrial lens. For more information about the ABViP CTP, please click on the following [link](#).

A webinar will be held on Thu 13<sup>th</sup> January 2022 17.00 – 18.30 (GMT) which will introduce the ABViP Programme, introduce each of the projects and provide an opportunity to have your questions answered. The final 30 minutes of the webinar will be an opportunity for potential applicants to meet with current doctoral students at UCL and University of Oxford. To register for this webinar, please [click here](#).

### **About the Department**

The Radcliffe Department of Medicine (RDM) is a large multi-disciplinary department that aims to tackle some of the world's biggest health challenges by integrating innovative basic biology with cutting edge clinical research.

We are made up of 5 smaller Divisions with internationally renowned programmes in areas such as cardiovascular medicine, diabetes and endocrinology, immunology, haematology and pathology. The Nuffield Division of Clinical Laboratory Sciences (NDCLS) is based at the John Radcliffe Hospital and studies the pathology of human disease from the molecular to the systems level; and aims to translate these findings into treatments for human disorders.

We also provide training to medical students, trainees and fellows in the area of clinical laboratory science through internationally acclaimed training programmes, bringing together all the clinical laboratory-based disciplines within the Oxford Medical School, including: microbiology, genetics, cellular pathology, haematology and clinical biochemistry.

### **About Oxford Biomedica**

Oxford Biomedica (OXB) is a pioneer of gene and cell therapy with a leading position in viral vector research and bioprocessing. Our mission is to deliver life-changing gene therapies to patients. OXB is an innovation and science focussed company which has developed a leading platform of novel technologies and capabilities. The OXB team provide design, development, bioprocessing and analytical development for gene-based medicines based on viral vectors, both for in-house products and for those developed with partner organisations. OXB has contract development and manufacturing organisation (CDMO) capabilities that support development of novel gene based medicines through all phases of clinical development to commercial manufacture. At Oxford Biomedica, we drive credible science to realise incredible results.

### **Entry requirements**

A UK Master's degree, or a minimum of an upper second-class UK Bachelor's degree, in a relevant discipline, or an overseas qualification of an equivalent standard. We particularly welcome applicants from disadvantaged backgrounds, or via an unconventional career path. If you're unclear as to whether you are eligible we would encourage you to submit an application regardless. You can also contact the project supervisor (see details below). To learn more about the policies in relation to diversity and inclusion at Oxford, please [click here](#) for further information.

Informal enquiries about the project should be addressed to [steve.hyde@ndcls.ox.ac.uk](mailto:steve.hyde@ndcls.ox.ac.uk)

### **Funding**

This BBSRC CTP ABViP Studentship is available to UK and Overseas (including EU) students. Full maintenance (stipend & fees) is available to UK and Overseas students for the duration of the four-year PhD. Note that up to a maximum of one fully-funded studentship allocations is available for Overseas students across the programme. The annual tax-free stipend for the PhD studentship is £16,077 (estimated).

### **English language requirements**

If your education has not been conducted in the English language, you will be expected to demonstrate evidence of an adequate level of English proficiency. The English language level for this programme is: **Standard**

### **Deadline and Application Process**

The deadline for submission is 12:00 midday on Fri 21<sup>st</sup> Jan 2022

To apply for this PhD studentship, you must submit a formal application to the DPhil in Advanced Bioscience of Viral Products course (Course code RD\_NG1) through UOXFs application portal by the above deadline. More information about the course and application process is available here: <https://www.ox.ac.uk/admissions/graduate/courses/dphil-advanced-bioscience-of-viral-products>