

## EPSRC Centre for Doctoral Training in Aerosol Science

**PhD Title:** Evaluation of the health impacts of aircraft nanoparticles using a surrogate soot source and in vitro cell exposure

### Details of Mentoring Team

|                  | Name                                      | Institution/Organisation                              | School/Department   |
|------------------|---|---|---|
| Lead academic(s) | Marc Stettler                             | Imperial College London                               | Civil and Environmental Engineering   |
| Co-supervisor(s) | Prof Peter Lindstedt<br>Prof Terry Tetley | Imperial College London<br>Imperial College London    | Mechanical Engineering<br>NHLI  |
| Partner(s)       | Dr Paul Williams<br><br>Dr Rachel Smith   | University of Manchester<br><br>Public Health England | Earth and Environmental Science<br><br>Nanoparticle Inhalation Research Group |

Is this a Tier 3 (industry led) proposal?                      Yes / No

### PhD Project Description

Context: Air travel is forecast to grow by 3.5% per year over the next two decades, leading to a doubling in passenger numbers from today's levels. Aircraft engines emit pollutants that degrade the air quality around airports, which may affect the health of local residents and employees at the airport. Specifically, nanoparticles emitted from gas turbine engines comprise soot aggregates, sulphur compounds, semi-volatile organic carbon and metallic ash. As the aircraft engine jet cools and mixes with the surrounding atmosphere, nucleation and condensation processes lead to a rapid growth in nucleation mode particles and condensation of semi-volatile compounds onto non-volatile soot aggregates. Aircraft nanoparticle emissions are characterised by their small size (<60 nm) relative to other combustion sources, e.g. road transport. They may therefore have particular health consequences that are currently under-studied and not well understood. A recent study<sup>1</sup> has conducted in-vitro toxicology tests to evaluate the effects on human bronchial epithelial cells to exposure to aircraft soot emissions from one type of aircraft engine. The authors found that the specific cellular response (cytotoxicity and oxidative stress) is dependent on the size and morphology of the soot particles, and fuel composition. However, much more work is needed to understand the different effects of aircraft nanoparticles and the role of their different properties and chemical constituents.

Specifically, the health impacts of aircraft nanoparticles may depend on the following properties:

- The size of the soot aggregates
- Their morphology and surface area
- Composition of coatings of organic carbon derived from unburned fuel or lubrication oil, e.g. presence of aromatic hydrocarbons
- Sulphur derived from naturally occurring sulphur in fuels
- Atmospheric ageing and oxidation of aerosols

Aims and objectives: This project aims to contribute to the understanding of the health impacts pathways of nanoparticles by developing and experimental methodology to test cellular responses to different sized nanoparticles and chemical compositions. The specific objectives of the project are to:

1. Review the literature on aircraft nanoparticle emissions, nanoparticle measurements near airports, and health impacts of different types and sources of combustion aerosols, including cellular response studies.
2. Develop laboratory source of soot particles that can be used to generate a tuneable surrogate aircraft aerosol in terms of particle size distribution and morphology, chemical composition and different coatings, and atmospheric ageing and oxidation. The soot source will be based on a burner design that has been used as a surrogate for aircraft soot particles before, and additional features such as organic carbon coatings and atmospheric ageing will be developed.
3. Use the surrogate soot source to design and conduct a comprehensive matrix of experiments to evaluate cellular responses to particle size and morphology, chemical composition of coatings, and atmospheric ageing. For example, aerosol classifiers could separate particular particle sizes before deposition on the cell culture to control for the effects of particle size. Similarly, particles could be coated with different materials, or have their coating removed by the use of a catalytic stripper. These methods are already established and involve exposure of primary human lung cells to increasing concentrations of the particles and determination of cell viability (MTT/LDH/apoptosis assays), mediator production (ie pro-inflammatory responses; ELISA assays), oxidative stress (ROS measurement), mitochondrial integrity (mitotracker), cellular antioxidant depletion (GSH oxidation) and particle uptake. Positive controls (eg ZnO nanoparticles) with known toxicity will be tested in parallel.
4. Evaluate potential health impacts of aircraft nanoparticle emissions around airports using existing evidence and new modelling works to understand how health risks may vary with distance from the airport, and other factors such as weather conditions.

Finally, new regulations on (non-volatile) particle emissions from aircraft engines means that there is a need to develop methods to calibrate particle analysers and particle sampling losses. The thematic broadening sabbatical will therefore explore the use of a surrogate soot source for instrument calibration.

References:

1. Jonsdottir, H.R., Delaval, M., Leni, Z., Keller, A., Brem, B.T., Siegerist, F., Schönerberger, D., Durdina, L., Elser, M., Burtcher, H., Liati, A., Geiser, M., 2019. Non-volatile particle emissions from aircraft turbine engines at ground-idle induce oxidative stress in bronchial cells. *Commun. Biol.* 2. doi:10.1038/s42003-019-0332-7
2. Stettler, M.E.J., Swanson, J.J., Barrett, S.R.H., Boies, A.M., 2013. Updated Correlation Between Aircraft Smoke Number and Black Carbon Concentration. *Aerosol Sci. Technol.* 47, 1205–1214. doi:10.1080/02786826.2013.829908

Thematic Area (see note below): Aerosols and Health

**Description of Thematic Broadening Sabbatical**

The thematic broadening sabbatical project will take place with Dr Paul Williams at University of Manchester. The project will be based on atmospheric aerosol measurements from the new urban supersite for monitoring ambient air quality. The student will complete relevant training and be given access to a whole range of ambient aerosol measurements. The project will analyse a year's worth of ambient air quality measurements and then focus in on specific pollution events. The project will emphasise the importance of meteorology in influencing properties of ambient aerosols and give an appreciation of how combustion aerosols are transported and processed in the atmosphere.

Thematic Area (see note below): Atmospheric Aerosol Studies

### Placement with Partner

The student will have the opportunity to spend time with the Nanoparticle Inhalation Research Group at Public Health England (information below). During the placement (to take place in year 3), the student will use PHE's air-liquid aerosol exposure systems (Vitrocell and/or Cultex) to expose cell cultures to a panel of particles produced by the surrogate soot source and to evaluate the effects of aerosol deposition. The student will also have the opportunity to interact with members of the Air Quality group at PHE, who provide advice and guidance in relation to air pollution.

The Nanoparticle Inhalation Research Group within the Toxicology Department at the PHE Centre for Radiation, Chemical and Environmental Hazards was established in 2009 to address concerns about the potential public health implications of the growing use of nanomaterials. The group undertakes basic research into the toxicity of nanomaterials and also potential exposure levels, with a focus on the inhalation pathway, and provides advice to Government (e.g. Defra) and other bodies, local, national and international, in this area. The group has state of the art facilities for the creation, delivery and characterisation of aerosols and associated chemical analysis facilities (e.g. ICP-MS/spICP-MS/Laser Ablation ICP-MS). We have undertaken toxicity studies for a range of nanomaterials including: silver, cerium dioxide, zinc oxide, iron oxides and carbon nanotubes. The key research interests of the group are: the deposition, clearance and translocation (toxicokinetics) of inhaled poorly soluble nanoparticles; the development and use of 'alternative' (in vitro) toxicity testing approaches using aerosol exposure air liquid interface (AE-ALI) systems; the use of radioactively labelled nanomaterials; the application of microarray techniques to evaluate gene expression changes in pulmonary cell types; the effect of inhaled nanomaterials and other air pollution components on pre-existing health conditions (e.g. asthma); the development of methodologies and approaches for on-line characterisation of nanoparticle aerosols, especially carbon nanotubes; and the chemical and biophysical nature of interactions between components of lung surfactant and nanoparticles.

### Explanatory Notes:

- Academic mentoring team: The lead academic supervisor and academic co-supervisor should be researchers in complementary thematic areas of aerosol science. The lead academic will be the main PhD supervisor; the co-supervisor will host the student in their research team for a 3-month thematic broadening sabbatical in the spring/summer of Year 1. There are no restrictions on the institution of the co-supervisor: they may be in the same institution as the lead academic or in one of the partnering institutions. It should be remembered that students will reside in Bristol in months 1-6 of Year 1. Many students may then prefer to move to the city they will reside in for the remainder of their PhD, so co-supervisors from nearby institutions (e.g. Leeds with Manchester, Bath with Bristol etc.) may be preferred by students, making recruitment easier. See Annex 1 for a full list of academic members of the supervisory team to assist lead academics in identifying an appropriate co-supervisor.
- Partner mentor: For *regular (non-Tier 3) studentships*, the partner will, most likely, have a broad interest in the area of the PhD project, but the research will be academic led. The partner will host the student for a 2-3 month placement at their organisation in Year 2 or 3. For *Tier 3 studentships*, the partner will co-fund the studentship at a level of £40k (£10k/year) and will work with the academic team to design a PhD project that addresses research of particular value to the partner.

- **Identifying partners for the second cohort:** Although a partner may be readily identified by the academic team for some projects, an appropriate partner may be less clear for others. If a partner cannot be readily identified by the academic team, the CDT leadership team can assist in identifying an appropriate partner to join the mentoring team. A list of industrial and public sector partners is given in Annex 2 to assist the academic team in identifying potential partners.
- **Thematic broadening sabbatical:** Supervised by the co-supervisor, this should have a focus in an area of research complementary to the final PhD project. Run as a mini-project during months 7-10 of the first year, a short project report is one element discussed in the progression interview at the end of Year 1. Brief project details should be provided on the form. Travel costs (e.g. travel to Leeds if the student is residing in Manchester, short term accommodation if further afield) and project costs will be paid by the CDT.
- **Placement with partner:** For regular studentships, this should be a 2-3 month placement anytime during Year 2 or 3. Placements for Tier 3 can be more extensive/frequent, as agreed by the mentoring team. A general overview of the likely area of activity should be provided, including potential training opportunities. Travel costs will be paid by the CDT.
- **Thematic Area:** You should identify the thematic area of the primary PhD project and the thematic broadening sabbatical. These should be in complementary areas of aerosol science using the definitions provided by the European Aerosol Assembly. The five thematic areas are: Basic Aerosol Processes, Aerosols and Health, Aerosol Technology, Atmospheric Aerosol Studies, Aerosol Measurement Techniques. See the "Working Groups" link at <http://www.gaef.de/ea/>. Also, note that further information on the coverage of the 5 themes is given at the links in the top left of the "Working Groups" page.

### More Information and How to Apply

Candidates who aspire to work in a multidisciplinary field, and hold or will achieve a minimum of an upper second-class undergraduate degree in any of these areas are encouraged to apply: chemistry, physics, biological sciences, life and medical sciences, mathematics and computer science, chemical and mechanical engineering, pharmaceutical and environmental sciences.

Visit our website:

<https://www.aerosol-cdt.ac.uk/>

Contact us:

[aerosol-science@bristol.ac.uk](mailto:aerosol-science@bristol.ac.uk)

Apply by 3rd February 2020– applicants with a suitable academic background will be invited to attend a recruitment and assessment day in Bristol on February 10<sup>th</sup>. Applications after this date will be subject to remaining availability of studentships.

### Equality, Diversity and Inclusion

We are committed to furthering issues of equality, diversity and inclusion and are keen to attract the most highly talented individuals from diverse backgrounds. The needs of individuals will be accommodated during the recruitment process and while studying with the CDT. Further information on our commitment to equality and diversity can be found on our website.

### Application Enquiries

Name: Kate Lucas (Aerosol CDT Administrator)

Email: [aerosol-science@bristol.ac.uk](mailto:aerosol-science@bristol.ac.uk)

Applications Tel: +44(0)117 928 8681

Applications WWW link: <https://www.aerosol-cdt.ac.uk/how-to-apply/>

## Annex 1: Core supervision team for EPSRC Centre for Doctoral Training in Aerosol Science

| Name              | Position                           | School/Department and Institution                               | Expertise   |
|-------------------|------------------------------------|---|---|
| Jonathan Reid     | Professor of Physical Chemistry    | Chemistry, University of Bristol                                | Aerosol microphysics, optical properties of aerosols, drug delivery to the lungs, formulation science, bioaerosol           |
| Bryan Bzdek       | Independent Research Fellow        | Chemistry, University of Bristol                                | Atmospheric aerosols, analytical techniques, new particle formation   |
| Michael Cotterell | Independent Research Fellow        | Chemistry, University of Bristol                                | Optical properties of aerosols, laboratory and airborne spectroscopy techniques   |
| Alberto Gambaruto | Lecturer                           | Mechanical Engineering, University of Bristol                   | Inhalation models and therapeutics  |
| Anwar Khan        | Senior Research Associate          | Chemistry, University of Bristol                                | Dispersion models, secondary organic aerosol, gas-particle partitioning   |
| Andrew Orr-Ewing  | Professor of Physical Chemistry    | Chemistry, University of Bristol                                | Optical properties of aerosols and spectroscopy   |
| Matt Rigby        | Reader in Atmospheric Chemistry    | Chemistry, University of Bristol                                | Atmospheric box models, dispersion models, secondary organic aerosol, gas-particle partitioning, machine learning, big data |
| Dudley Shallcross | Professor of Atmospheric Chemistry | Chemistry, University of Bristol                                | Atmospheric Chemistry research group, Atmospheric box models  |
| Matt Watson       | Reader                             | Earth Sciences, University of Bristol                           | Atmospheric aerosols, instrumentation, geo-engineering  |
| Michael Hill      | Professor of Inorganic Chemistry   | Chemistry, University of Bath                                   | Aerosol-assisted chemical vapour deposition   |
| Andrew Johnson    | Lecturer                           | Chemistry, University of Bath                                   | Aerosol-assisted chemical vapour deposition   |
| Matthew Jones     | Lecturer                           | Pharmacy and Pharmacology, University of Bath                   | Inhalation therapeutics and drug delivery (esp. device usability), formulation science                                      |
| Anton Souslov     | Lecturer                           | Physics, University of Bath                                     | Levitation in air. Theory and modelling of soft materials. Phases of soft matter.   |
| Adam Squires      | Senior Lecturer                    | Chemistry, University of Bath                                   | Basic process studies and new techniques for aerosols (e.g. synchrotron small-angle X-ray)                                  |
| Jethro Akroyd     | Senior Research Associate          | Chemical Engineering and Biotechnology, University of Cambridge | Combustion, nanoparticles, soot   |

| <b>Name</b>        | <b>Position</b>                 | <b>School/Department and Institution</b>                        | <b>Expertise</b>   |
|--------------------|---------------------------------|---|--|
| Adam Boies         | Director of ANAM, Reader        | Engineering, University of Cambridge                            | Particle synthesis using aerosol flow reactors (e.g. carbon nanotubes, carbon black)   |
| Megan Davies Wykes | Lecturer in Engineering         | Engineering and Applied Mathematics, University of Cambridge    | Pollution dispersion modelling, experimental fluid dynamics  |
| Simone Hochgreb    | Professor of Engineering        | Engineering, University of Cambridge                            | Combustion, particle formation and optical diagnostics   |
| Markus Kraft       | Professor of Engineering        | Engineering, University of Cambridge                            | Particle synthesis using aerosol flow reactors (e.g. carbon nanotubes, carbon black)   |
| Sebastian Mosbach  | Senior Research Associate       | Chemical Engineering and Biotechnology, University of Cambridge | Computational modelling  |
| Robert Chilcott    | Professor in Pharmaceutics      | Life and Medical Sciences, University of Hertfordshire          | Aerosol dispersal, exposure to chemical and radiological aerosols, toxicology and exposure remediation                             |
| Richard Greenaway  | Senior Research Fellow          | Physics, University of Hertfordshire                            | Particle instrumentation, data-analysis, optics  |
| Ian Johnston       | Associate Professor of Research | Engineering, University of Hertfordshire                        | Microfluidic systems, experimental aerosol sampling and collection techniques, biodetection  |
| Joanne Larnier     | Senior Research Fellow          | Life and Medical Sciences, University of Hertfordshire          | Chemical analysis of aerosol composition, bio exposure   |
| Daniel McCluskey   | Associate Professor of Research | Engineering, University of Hertfordshire                        | Aerosol sampling techniques & system design, fluid dynamics, advanced microfluidics, biodetection                                  |
| Darragh Murnane    | Professor in Pharmaceutics      | Life and Medical Sciences, University of Hertfordshire          | Inhalation therapeutics and particle-lung interactions, delivery device design   |
| Chris Stopford     | Associate Professor             | Physics, University of Hertfordshire                            | Optical properties of aerosol and novel instrumentation  |
| Laura Urbano       | Lecturer in Pharmaceutics       | Life and Medical Sciences, University of Hertfordshire          | Novel nano-diagnostics and therapeutics. In-vitro models to contribute to the fields of drug safety, drug delivery and translation |
| Denis Doorly       | Professor of Fluid Mechanics    | Engineering, Department of Aeronautics, Imperial College London | Fluid mechanics  |

| <b>Name</b>       | <b>Position</b>                                      | <b>School/Department and Institution</b>                     | <b>Expertise</b>   |
|-------------------|--|--|--|
| Yannis Hardalupas | Professor of Multiphase Flows                        | Mechanical Engineering, Imperial College London              | Multiphase flows and sprays, droplet and nanoparticle sizing, exhaust after-treatment, combustion in gas turbines and IC engines |
| Jerry Heng        | Reader in Particle Technology                        | Chemical Engineering, Imperial College London                | Nucleation, crystallisation, surface properties, powders   |
| Peter Lindstedt   | Professor of Thermofluids                            | Mechanical Engineering, Imperial College London              | Combustion and soot  |
| Asha Patel        | Lecturer in Cell & Gene Therapy                      | National Heart & Lung Institute, Imperial College London     | Inhaled delivery of advanced therapeutics such as mRNA   |
| Alex Porter       | Professor of Bio-imaging and Analysis                | Materials, Imperial College London                           | Toxicology of particles, enhanced Electron Microscopic Imaging   |
| Marc Stettler     | Lecturer in Transport and Environment                | Civil and Environmental Engineering, Imperial College London | Nanoparticle emissions sources, measurements and models, transport sector, ambient pollution                                     |
| Terry Tetley      | Professor of Lung Cell Biology                       | National Heart & Lung Institute, Imperial College London     | Cellular and molecular mechanisms of pulmonary diseases  |
| Omar Usmani       | Reader in Respiratory Medicine                       | National Heart & Lung Institute, Imperial College London     | Inhalation therapeutics, drug delivery to the lungs  |
| Andrew Bayly      | Professor of Chemical Engineering                    | Chemical and Process Engineering, University of Leeds        | Materials processing, spray drying and particle fabrication  |
| Dwayne Heard      | Professor of Physical Chemistry                      | Chemistry, University of Leeds                               | Laboratory studies of atmospheric aerosols and oxidation chemistry   |
| Nik Kapur         | Professor of Applied Fluid Mechanics                 | Mechanical Engineering, University of Leeds                  | Droplets, sprays and coatings, fluid flow and heat transfer  |
| Ben Murray        | Professor of Atmospheric Science                     | Earth and Environment, University of Leeds                   | Laboratory studies of atmospheric aerosols including ice   |
| Catherine Noakes  | Professor of Environmental Engineering for Buildings | Civil Engineering, University of Leeds                       | Indoor air quality and airborne infection, ventilation and building performance  |
| John Plane        | Professor of Physical Chemistry                      | Chemistry, University of Leeds                               | Laboratory studies of interstellar chemistry on dust   |
| Andrew Brass      | Professor of Bioinformatics                          | Computer Science, University of Manchester                   | Health informatics and immunology, allergies and environmental factors   |

| <b>Name</b>        | <b>Position</b>  | <b>School/Department and Institution</b>                              | <b>Expertise</b>   |
|--------------------|--|---|--|
| Hugh Coe           | Professor of Atmospheric Composition                   | School of Earth and Environmental Sciences, University of Manchester  | Particle detection, physical and chemical characterization, gas/particle conversion, instrumentation, sampling and data analysis methods |
| Sheena Cruickshank | Professor of Public Engagement and Biomedical Sciences | Biological Sciences, University of Manchester                         | Impact of pollutants and airborne microbes on barrier cells, public engagement and citizen science                                       |
| Amanda Lea-Langton | Lecturer in Bioenergy Engineering                      | Mechanical, Aerospace and Civil Engineering, University of Manchester | Smoke formation and organic carbon emissions, environmental aerosol  |
| Gordon McFiggans   | Professor of Multiphase Atmospheric Processes          | School of Earth and Environmental Sciences, University of Manchester  | Atmospheric Composition, Clouds & Aerosols, Computer Modelling, Marine Atmospheric Chemistry   |
| Benedict Rogers    | Reader   | School of Mechanical, Aerospace and Civil Engineering                 | Smoothed particle hydrodynamics (SPH) with novel applications to aerosol problems  |
| David Topping      | Senior Lecturer  | Earth and Environmental Science, University of Manchester             | Aerosol microphysics, air quality, environmental aerosol, computer modelling and data science/ machine learning                          |
| Paul Williams      | Independent Research Fellow                            | Earth and Environmental Science, University of Manchester             | Vehicle emissions, new instrumentation, environmental aerosols   |

## **Annex 2: Current List of Potential Partners**

3M Healthcare UK Ltd  
Agilent  
Alphasense  
Astra Zeneca  
Bayer Crop Science - Bioaerosol detection  
Bespak Europe Ltd  
Biral  
Cambustion  
Centre for Ecology and Hydrology  
Chiesi  
CMCL Innovations  
CN Bio  
DEFRA  
Droplet Measurement Technologies  
DSTL  
Dyson  
Echion Technologies  
Emissions analytics  
Environment Agency  
Filter Integrity  
GSK  
Intertek Melbourn  
Johnson Matthey  
LettUs Grow  
Malvern Instruments  
MedPharm  
Met Office  
NanoPharm  
National Physical Laboratory  
Nyquist Solutions Ltd  
Philips / PMO and Technology Respiratory Drug Delivery  
Pirbright  
Public Health England  
Rolls Royce  
Rothamsted Research  
RSK  
Siemens  
Steer Energy  
Syngenta  
TH Collaborative Innovation  
Tortech  
Trolex  
TSI  
Vectura  
Venator  
Waters Mass Spec