

## Editorial

For the second time in succession, AIR<sup>2</sup> highlights two exposure platforms in this 16<sup>th</sup> issue. Although exposure platforms are the cornerstone of most radiation protection research activities, featuring two together also indicates our lack of knowledge and visibility of many of the databases, sample banks and cohorts available to our research community. These less visible infrastructures are also the most fragile. Frequently created during a European project, they fall dormant afterwards through lack of sustainable funding, when they could have been so useful for future research if kept active and updated. But AIR<sup>2</sup> could help to keep them under the spotlight. The more attractive they are, the greater their chances of attracting new users. AIR<sup>2</sup> aims to optimize all efforts to revive these databases, sample banks and cohorts.

**Dr Laure Sabatier, CEA**

## The floor to...

The goal of task 6.3, based on previous experience gained from DOREMI and OPERRA and the results of tasks 6.1 and 6.2, is to facilitate access to all suitable infrastructures identified and, in some cases, to rare or unique infrastructures for radiation protection research.

For its two infrastructure calls, CONCERT has chosen to take an open-minded approach and to accept all the various types of infrastructures, on condition that the cost of the activities

for the submitted projects is included in the global cost of the research project.

The funding scheme creates access to the infrastructures but does not guarantee their sustainability nor any potential future access for our community.

The work of task 6.3 involves identifying the support needs to achieve sustainability. These include the necessary maintenance and upgrade of materials, maintaining skills and state-of-the-art facilities, developing specific training courses for relevant infrastructures, increasing accessibility through use of simple procedures and common protocols, and finally guaranteeing

the quality of the results produced. These needs are all components of the strategy to ensure the continued existence of the infrastructures and thus of the research programmes related to the priorities defined in the SRA developed by the platforms MELODI, ALLIANCE, NERIS, EURADOS, and more recently EURAMED.

In summary: WP6 contributes to establishing links between infrastructures and researchers; it provides the information needed to develop opportunities with

the possibility of infrastructure access and funding; it builds and offers possible ways to implement best practice and it focuses on synergies. The true success of WP6 will be measured when a roadmap approved by all parties is implemented in order to increase access to and use of these infrastructures.

### Strategy for Facilitating Access to infrastructures

**Jean-Michel Dolo**  
CEA  
CONCERT WP6.3



Photo: J-M Dolo/CEA

Issue **16**  
April 2017

 **CONCERT  
ORNER**

### Future events:

2nd Call

1 Mar 2017: Opening date

May 2<sup>nd</sup>, 2017: Deadline

### WP 6 News:

Next WP6 meeting:

May 22<sup>nd</sup>, Budapest, Hungary

### AIR<sup>2</sup>D<sup>2</sup>:

- Please complete the online [form\(s\)](#) to register your infrastructure(s) in the database.
- A new option to feature your infrastructure is now available: [add document](#).

### Contents:

Exposure platforms [Changing dose rate \(SU\)](#)

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Analytical platforms, Models, Tools [Advanced Technologies Network Center](#)

### Next issue

May 2017



## Changing dose rate exposure facility

Exposure of cells to continuously changing photon dose rate

**E**xposure scenarios where the dose rate is continually changing are very common. A good example is aircraft flight where the dose rate of cosmic radiation can change 16-fold during take-off and landing. Moreover, there are many accidental exposure scenarios where either the sources or the exposed subjects are in motion with respect to one another. Despite the fact that many exposures involve changing dose rates, the vast majority of research studying the effects of ionising radiation is performed exposing samples at constant dose rates. It is



The facility inside an incubator

The peristaltic pump is not visible. The X-ray tube is positioned below the incubator and the whole setup is enclosed in a lead cabinet for safe use

interconnected by a silicone tube via a peristaltic pump. Cell samples can be positioned on top of the tanks. The facility fits inside a 164 l cell incubator modified so that there are no wires or electronic components in its bottom plate. An X-ray tube is placed under the incubator and the distance from the X-ray source to the bottom of the facility is ~ 30 cm. The beam angle, as given by the manufacturer, is 40° x 55°.

During exposure, the pump transfers the shielding medium, an aqueous solution of barium chloride, from one tank (increasing dose-rate, IDR) into the other (decreasing dose-rate, DDR), resulting in an exponential, 14-fold dose-rate change during the exposure. Tank 3 (average dose-rate, ADR) contains a volume of barium chloride resulting in the same dose-rate on top of the tank as the average dose-rate on top of tanks 1 and 2. The exposure is monitored with an ionisation chamber positioned on the tank that is acting as the IDR tank, and terminated when the starting conditions have been reversed on top of tanks 1 and 2. Consequently, the same total dose will have been delivered on top of all three tanks when the exposure is terminated.

The facility makes it possible to characterise the cellular response to changing dose rates. The design and low building cost of the device permit users to customise and build a device to suit their particular needs, encouraging other research groups to contribute to the understanding of the effects of changing dose rates.



Andrzej Wojcik

Photo: Andrzej Wojcik (SU)



ID Card:

**Exposure type:**

External

**Source:**

X-ray machine

**Dose rate:**

2.2 to 37 mGy per minute

**Irradiation type:**

photons, vertical beam

**Irradiated organism type:**

Cells in culture

**Address:**

Stockholm University

**Access:**

free, decision by source owner

**Supporting lab:**

Biomolecular and cell culture lab

**Internet link:**

**Contact:**

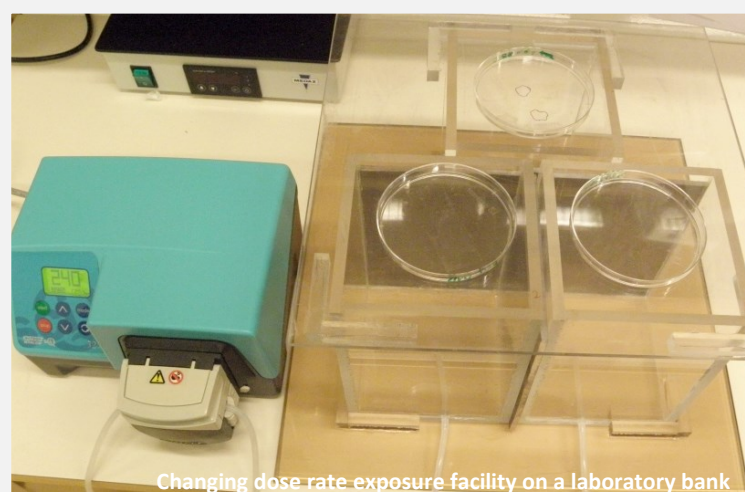
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**Related to:**

MELODI, EURADOS, RENEB



Changing dose rate exposure facility on a laboratory bank

Petri dishes are placed on top of tanks between which a barium chloride solution is pumped with the help of a peristaltic pump (green). A third tank (visible behind the two front tanks) is permanently filled with a volume of barium chloride that yields the average dose rate. Consequently, cells on all three tanks receive the same dose

Photo: Andrzej Wojcik (SU)

possible that the technical limits of the irradiation equipment used may prevent other types of exposure scenarios. However, effects from such exposures may be highly relevant for the assessment of radiation risk. Thus, it is surprising that research on the biological effects of changing dose rates has, until recently, been neglected.

To study the effects of changing dose rates, we have constructed a facility where three samples can be simultaneously irradiated with X-rays either at an increasing, a decreasing, or a constant dose rate. The facility fits inside a 37°C incubator that can be positioned above an X-ray tube or a gamma source. Cells in tubes, flasks or Petri dishes can be simultaneously exposed to an increasing, a decreasing and a constant dose rate in the range of 2.2 to 37 mGy per minute.

The facility is composed of three identical Plexiglas tanks, separated by 4 mm lead plates to absorb scattered radiation. Tanks 1 and 2 are



# Exposure platforms

## Low dose rate facility at Stockholm University

### Low dose rate exposure facilities for cells and animals

**S**tockholm University was founded in 1878. Today, it has 70,000 students, 1,800 doctoral students and a staff of 5,000 who are active within science, the humanities, social sciences and law. The first Chair of Radiobiology was appointed in 1962, at a time when the work focused on genetics and plant breeding. In 1972, Radiation Biology moved to the Wallenberg Laboratory at the new campus in Frescati, and in 1985 to the Arrhenius Science Laboratories.

In the 1970's, low dose radiation facilities were constructed for field experiments, mainly for plant genetics and genotoxicology. At that time there was already a strong focus on DNA damage,

shielding to a few  $\mu\text{Gy/h}$ . Mice can be exposed chronically and exposure time should not exceed 4 weeks. The animals are hosted in standard cages with space for up to 5 mice per cage. The facility can accommodate four cages placed one on top of the other, providing a gradient of dose rates (picture 2). After exposure, the mice can be kept in the animal facility for extended periods depending on the choice of endpoints.

The construction has been approved by the Swedish Radiation Safety Authority (SSM) and complies with the ethical rules for animal experiments. The primary objective is to provide scientists in the field of radiation protection research with access to a low dose and dose rate exposure facility for short term exposure of mice.

The animal exposure facility is located in the Stockholm University animal facility, and includes animal care, animal exposure and post irradiation handling, for example, preparation of organs/samples at different times post irradiation. This radiation facility is primarily constructed for the study of biomarkers in response to low doses and dose rates and for studies of the mechanisms behind cellular/organ responses. It may also be used for pilot studies where only a small number of animals are needed.

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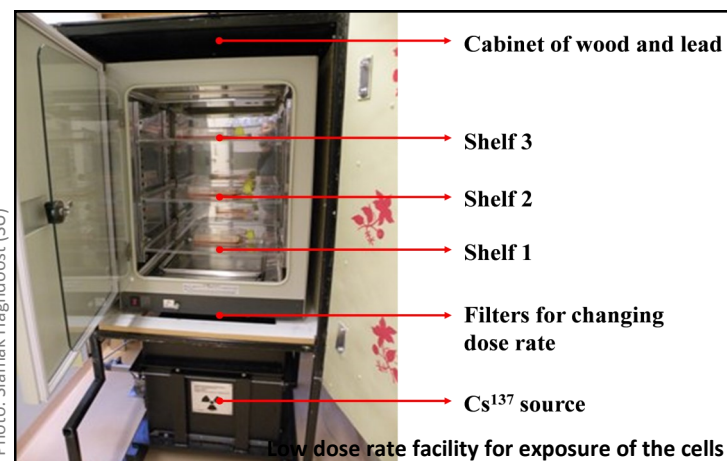


Photo: Siamak Haghdoust (SU)

and several new methods were invented to measure DNA strandbreaks as well as chromosomal damage in plants and eukaryotic cells.

In the last two decades, the interest of the research groups has gradually moved towards risk estimates of low doses and dose rates, and to exploration of new technologies such as omics to study the cellular responses to doses in the mGy range. Thus new radiation exposure facilities were needed in the department and, with the help of skilled technicians and an excellent workshop, several new facilities were constructed, as described below.

At present two radiation facilities with caesium sources are available for chronic exposure of cells in culture, with dose rates ranging from 1 mGy/h up to 50 mGy/h, and dose rate can be decreased by lead shielding to a few  $\mu\text{Gy/h}$  (picture 1) [1, 2]. A new radiation facility for animal exposure was constructed in 2015 and is equipped with a caesium source. This facility is constructed for exposure of mice to low doses and low dose rates. The dose rates range from 1 mGy/h up to 70 mGy/h and dose rate can be decreased by lead



**Siamak Haghdoust**

Photo: Siamak Haghdoust (SU)



#### ID Card:

##### Exposure type:

Low dose rates external gamma radiation

##### Source:

Cesium 137

##### Dose rates

##### Cell culture facility:

From 1 to 50 mGy/h with lead shielding

##### Animal facility:

From 1 up to 70 mGy/h with lead shielding

Housing capacity: 4 cages and 5 mice per cage

##### Preferred type of organism for irradiation:

Mouse

##### Exposure time:

Up to 4 weeks

##### Address:

Centre for Radiation Protection Research  
Department of Molecular Bioscience, Wenner-Gren Institute  
Stockholm University  
10691 Stockholm  
Sweden

##### Access:

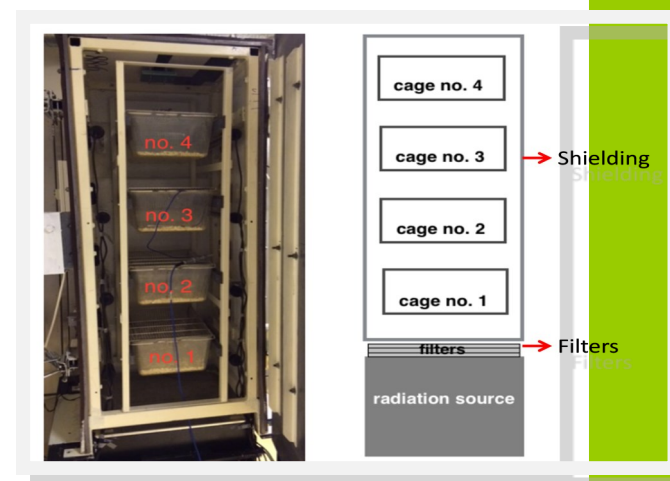
Joint research collaboration and upon ethical approval by the ethical committee

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##### Related to:

MELODI, EURADOS



**Low dose rate facility for animal exposure**

Photo: Siamak Haghdoust (SU)



## Advanced Technologies Network (ATeN) Center

A large research infrastructures for advanced biotechnologies

The Advanced Technologies Network (ATeN) Center, directed by Prof. Maurizio Leone, is a centre of excellence of the University of Palermo (Sicily) which provides cutting-edge research, development and service activities for technological transfer to the public and private sectors. The Center consists of three macro-areas (Cellular and Molecular Biotechnology, In vivo Analysis, Biocompatible Materials and Systems) in which scientists with different backgrounds (e.g. biotechnology,

of genetic and protein profiles and of molecular pathways, experimental cellular and animal models of disease; identification of specific response markers of cells and tissues to exposure to ionising radiation and/or molecules with biological activity and potential pharmacological activity; validation of products for molecular diagnostics; development of services for advanced diagnostics and for drug discovery; development of bioinformatics products (acquisition, storage, distribution, analysis and interpretation of the data mainly for molecular biology, genetics and biochemistry).

The In Vivo Analysis macro-area, with two enclosures containing small animals and zebrafish, carries out analyses on the effects of ionising radiation and the

testing of drugs, biomaterials, biomarkers and radiopharmaceuticals, as well as functional analyses for the production of primary cultures from transgenic organisms and 3D imaging.

Multiple bioimaging techniques are available to explore the biological structure and function of molecules in live cells and in tissues by means of 3D and 4D measurements.

Confocal and multiphoton microscopy, atomic force microscopy, together with advanced spectroscopy techniques (e.g. Raman, EPR, NMR) can be applied to analyse biological, physical and chemical phenomena in order to characterise the material properties.



Photo: Agenzia CMC Studio

**Maurizio Marrale**



Photo: Agenzia CMC Studio

**Ion PGM™ System for Next-Generation Sequencing**

biology, chemistry, physics, engineering, medicine, bioinformatics) work together to produce the technological know-how needed to achieve highly competitive scientific results. Due to its sophisticated structure and equipment (25 laboratories housed in 2500 m<sup>2</sup> with approx. 100 instrumentation facilities), ATeN is among the few centres in the world able to provide a production chain ranging from the synthesis of materials to in vivo tests.

The macro-area of Cellular and Molecular Biotechnologies deals with the production and propagation of stem cells and primary cell cultures, large-scale analysis of DNA, RNA and proteins. The laboratory of genomics and proteomics provides molecular analysis at advanced technological level. The laboratory works in different advanced sectors through the analysis of large families of genes, proteins, enzymes and metabolites. These sectors include: development and technological improvement of drugs including proteins, vaccines and monoclonal antibodies, which are largely obtained from targeted application of genetic modification techniques and personalised medicine; characterisation, through the analysis

**Laboratory of Pulsed Electron Paramagnetic Resonance**



Photo: Agenzia CMC Studio



### ID Card:

#### Analytical platform type:

Biological dosimetry and physical retrospective dosimetry, exposure markers, proteomics, genome sequencing, transcriptome sequencing, transcriptomics, metabolomics, exosomes, small molecules

#### Main techniques proposed:

Panel of cytogenetic assays, gene expression assay, protein markers, EPR/TL dosimetry, gamma spectrometry, microscopy

#### Capacity:

20 measurements per week

#### Waiting time:

None

#### Duration of experiment:

Dependent on experiment and assay

#### Address:

Viale delle Scienze Edificio  
18 I-91128 Palermo (Italy)

#### Access:

Free

#### Internet link:

<http://www.chab.center/home-en>

#### Contact:

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+39 091 23899073

#### Related to:

MELODI, EURADOS, RENEB

## Future events:

### CONCERT Short Courses

**24-28 April 2017**  
Emergency and recovery preparedness, and response  
NCCRP, Sofia, Bulgaria  
**Contact:**  
[n.chobanova@nccrp.org](mailto:n.chobanova@nccrp.org)

**24 April-5 May 2017**  
Cellular effects of ionising radiation – introduction to radiation biology  
Stockholm University, Sweden  
**Contact:**  
[andrzej.wojcik@su.se](mailto:andrzej.wojcik@su.se)

**22 May-2 June 2017**  
Modelling radiation effects from initial events  
University of Pavia, Italy  
**Contact:**  
[Andrea.Ottolenghi@unipv.it](mailto:Andrea.Ottolenghi@unipv.it)

**6-16 June 2017**  
Assessing risk to humans and the environment  
NMBU, Oslo, Norway  
**Contact:**  
[deborah.oughton@nmbu.no](mailto:deborah.oughton@nmbu.no)

### Other Events

**25-27 April 2017**  
**COMET final event**, Bruges, Belgium

**8-11 May 2017**  
**ConRad 2017**, Bundeswehr Institute of Radiobiology, Munich, Germany

**14-19 May 2017**  
**NEUDOS13** Neutron and Ion Dosimetry Symposium  
Krakow, Poland

**23-26 May 2017**  
**Opera final event**  
Budapest, Hungary

**11-17 June 2017**  
**RAD 2017** Fifth International Conference on Radiation and Applications in Various Fields of Research  
Budva, Montenegro

**3-8 September 2017**  
**ICRER 2017**, 4th International conference on Radioecology and Environmental Radioactivity,  
Berlin, Germany

**10-12 October 2017**  
**Joint ICRP-RPW 2017**  
Paris, France

**5-11 November 2017**  
**MICROS 2017**, 17<sup>th</sup> International Symposium on Microdosimetry,  
Venice, Italy

## Issue

## Exposure platforms

## Databases, Sample banks, Cohorts

## Analytical platforms Models & Tools

### Published to date:

Oct 2015, #1  
Nov 2015, #2

FIGARO

B3, Animal Contamination Facility

FREDERICA

The Wismut Cohort and Biobank

RENEB

The Hungarian Genomics Research Network

Dec 2015, #3  
Feb 2016, #4

Pulex Cosmic Silence

SNAKE

STORE

French Haemangioma Cohort and Biobank

METABOHUB

Dose Estimate, CABAS, NETA

Mar 2016, #5  
Apr 2016, #6

Radon exposure chamber  
Biological Irradiation Facility

3-Generations exposure study  
Wildlife TransferDatabase

PROFI  
Radiobiology and immunology platform (CTU-FBME)

May 2016, #7  
Jun 2016, #8

CIRIL  
Mixed alpha and X-ray exposure facility

Portuguese Tinea Capitis Cohort  
Elfe Cohort

LDRadStatsNet  
ERICA Tool

Jul 2016, #9  
Sep 2016, #10

SCRS-GIG  
Facility radionuclides availability, transfer and migration

RES<sup>3</sup>T  
INWORKS cohort

CROM-8  
France Génomique

Oct 2016 #11

LIBIS gamma low dose rate facility ISS

JANUS

Transcriptomics platform SCK CEN

Nov 2016, #12  
Dec 2016, #13

Microtron laboratory  
Nanoparticle Inhalation Facility

EPI-CT Scan cohort  
UEF Biobanking

CATI  
The Analytical Platform of the PREPARE project  
HZDR Radioanalytical Laboratories

Feb 2017, #14

Infrastructure for retrospective radon & thoron dosimetry

Chernobyl Tissue Bank

Mar 2017, #15

Alpha Particles Irradiator Calibration Laboratory at KIT

SYMBIOSE

Apr 2017, #16

Changing Dose rate (SU)  
Low dose rate (SU)

Advanced Technologies Network Center

### Coming soon:

May 2017, #17

To Be Announced

To Be Announced

To Be Announced