

Equivalent dose measurements of astronauts for ISS space missions between 2007-2016

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Outline

- Biodosimetry - Health Canada's space biodosimetry program and limitations
- Equivalent dose estimates and particle fluxes inside ISS for CSA and ESA missions from 2007 – 2016 with OLTARIS
 - GCR and TPs effects only
- Monte Carlo programs for cellular biology
- Conclusions and next steps

Biodosimetry

- **Method used to obtain dose estimates for individuals exposed to ionizing radiation based on their biological responses at the cellular level**

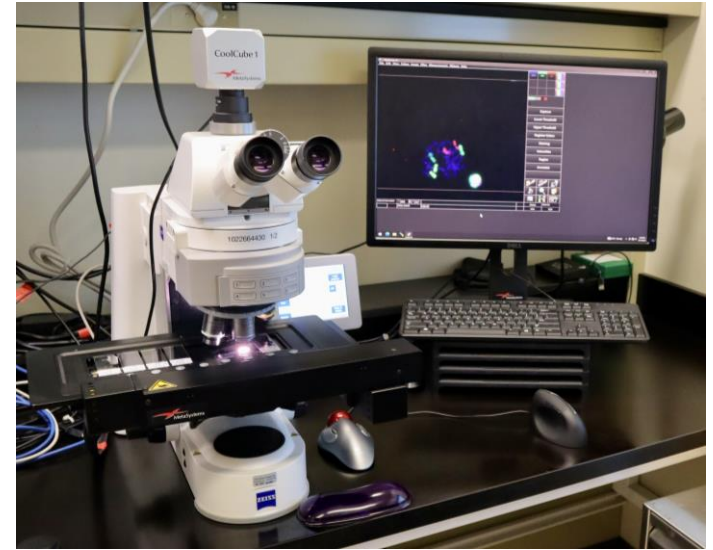
Useful in:

- Cases of accidental exposure
 - When physical dosimetry is called into question or not possible
 - Cases of nuclear emergencies
 - Determining astronauts' exposure to space radiation
-
- Our Division focuses on all these situations



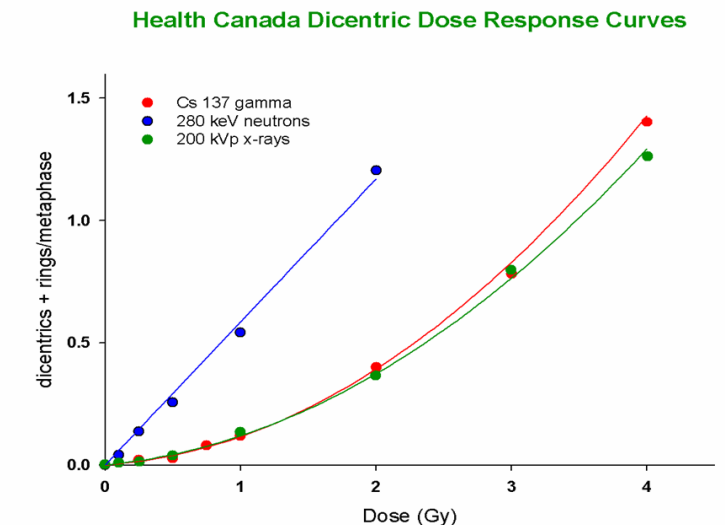
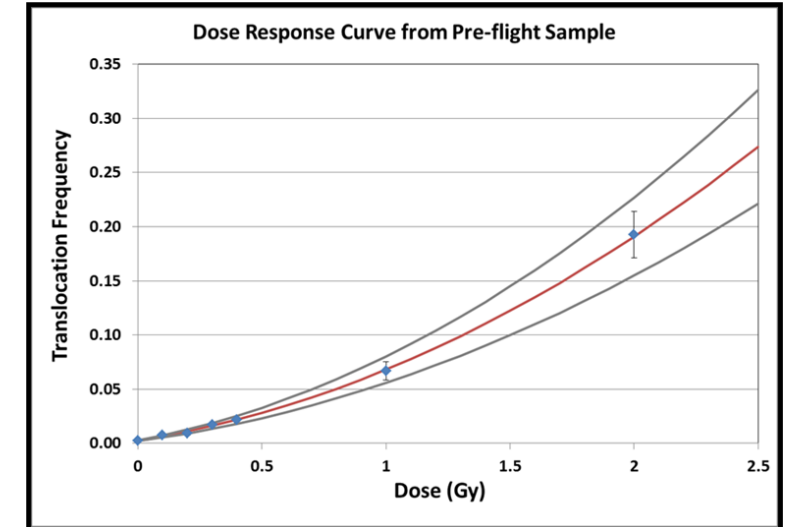
Biodosimetry

- At Health Canada, we are equipped to perform the following assays:
 - Translocation analysis by the Fluorescence *in situ* hybridization (FISH)
 - Dicentric Chromosome Assay (DCA)
 - Cytokinesis Block Micronucleus (CBMN)
 - $\gamma\text{H}_2\text{AX}$ assay
 - And more ...



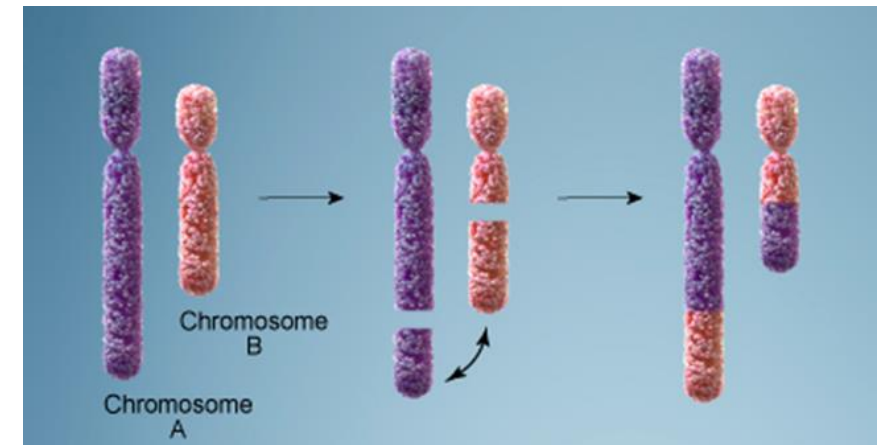
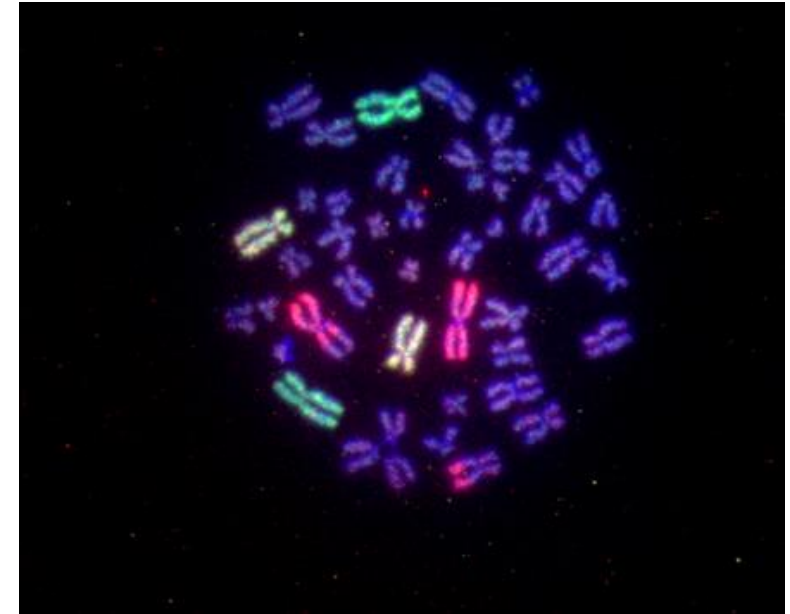
Biodosimetry

- All biodosimetry methods require the use of a **calibration curve**:
 - Mathematically models biological responses to irradiations at set doses
 - Used to obtain dose estimates
- Calibration curves have different shapes depending on the radiation source:
 - Linear for high LET radiation
 - Quadratic for low LET radiation
- The shape affects the accuracy of the dose estimates



Biodosimetry

- For astronauts, translocation analysis is our go-to biodosimetry assay
- **Measures stable translocations**
 - Occurs when chromosome segments are exchanged but no genetic information is lost
 - **Methodology** involves:
 - Irradiating blood in a 250kVp X-ray cabinet
 - Following various preparation steps, lymphocytes with painted chromosomes are scored by microscopy to measure frequency of stable translocations



Source: U.S. National Library of Medicine

Health Canada's space biodosimetry program

- Biodosimetry is currently a medical requirement for astronauts as specified in the ISS MED Vol. B Appendix A for CSA
- Career radiation dose limit for CSA astronauts is 1Sv
- Health Canada conducts biodosimetry analysis for CSA astronauts after every space flight
 - We produce personalized calibration curves for each astronaut
 - Our translocation analysis scoring system has been harmonized with NASA to ensure comparability of data
 - ESA analysis was also done by HC from 2007 until 2016 but has since been discontinued
- Overall, program has completed dose assessments on 11 astronauts

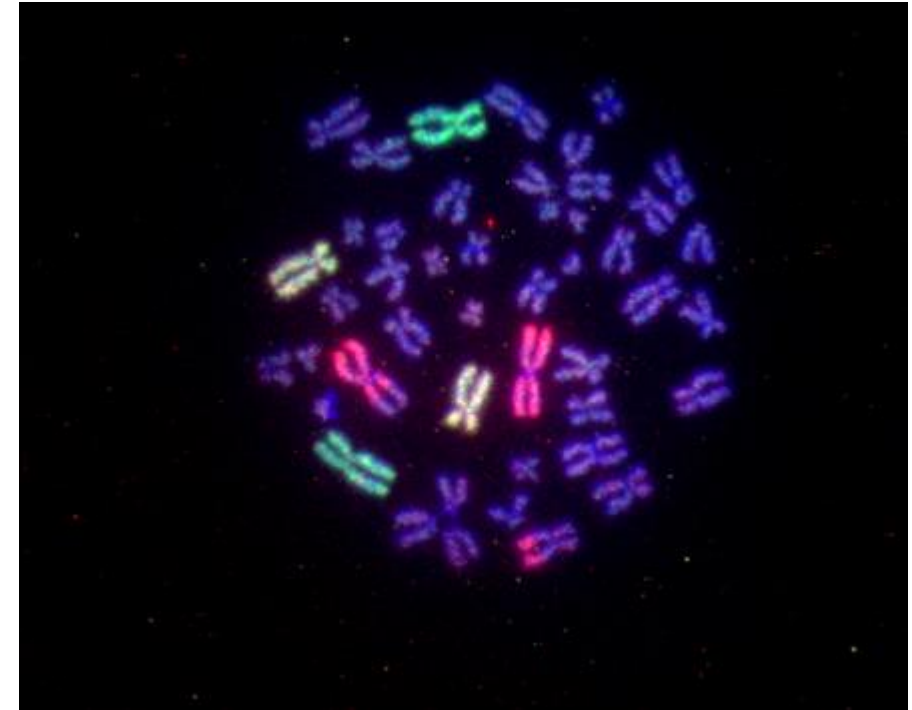
FISH procedure for astronauts

- ✓ Blood samples collected under the supervision of the flight surgeon pre- and post-flight
 - Pre-flight sample is used to establish a baseline
 - 34 mL of blood drawn 35-90 days before flight
 - Post-flight sample drawn immediately after flight and 6 months after
 - 17 mL of blood drawn 7-14 days post-flight
 - 17 mL blood drawn 6-18 months post-flight
- ✓ Samples are delivered within 24-48 hours after collection
- ✓ Dose estimates provided approximately 1 month after receiving the blood



FISH scoring methodology

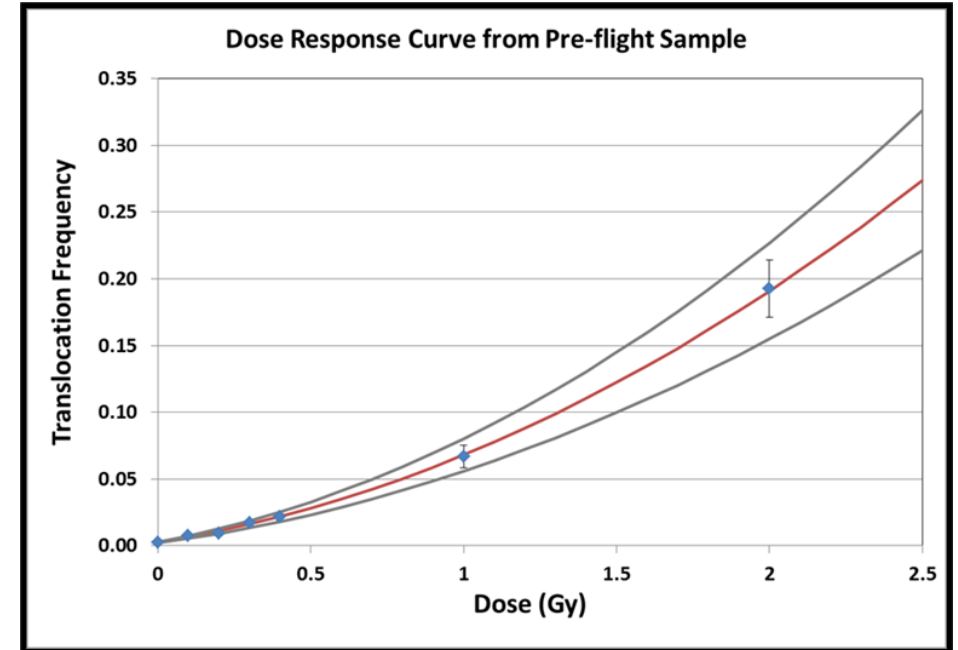
- We score 10,000 cells in metaphase pre-flight at 0 Gy to determine the background damage, then between 500 (high doses) – 4000 (low doses) to produce the calibration curve.
- 1,000 cells are scored in both post-flight draws.
 - Use our own damage scoring system that has been harmonized with NASA
 - Score different types of simple DNA damage including Apparently Stable Translocations (AST)
 - Also, take note of but not include complex damage (3 or more cuts in 2 or more chromosomes).



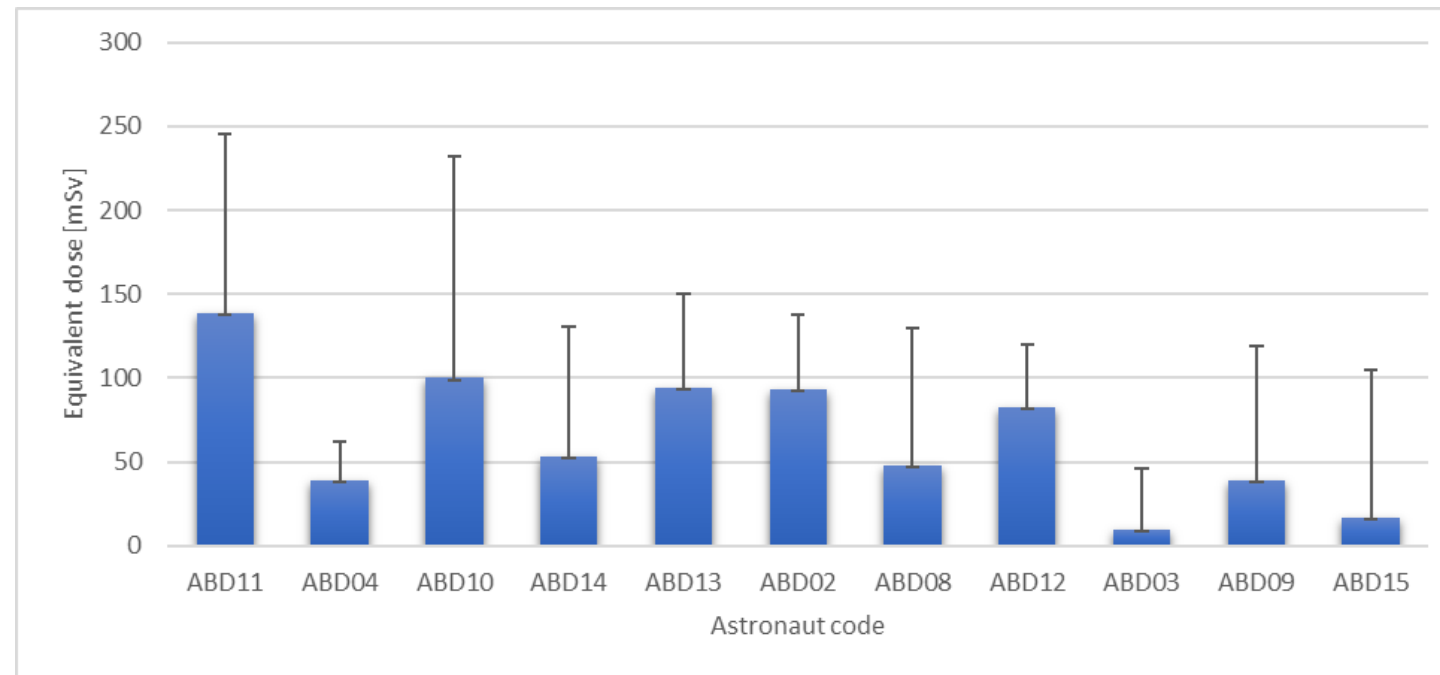
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FISH

- At HC we build personalized astronaut calibration curves by finding the translocation frequency (translocations/cells scored) for different doses
 - Post-flight (immediately and 6 months), translocation frequencies measured in blood samples and matched to these calibration curves to obtain dose estimates



Health Canada's space biodosimetry program: at a glance



Total: 11 individuals over 10 years

Mostly 6-month missions

Health Canada's space biodosimetry program: limitations

- Our calibration curves are limited to X-ray sources (due to laboratory restrictions)
 - To address this issue, we are exploring the use of Monte-Carlo based methods to create mixed source calibration curves that more accurately reflect space radiation environments
 - With these calibration curves, we seek to determine a correction factor to apply to our current assay

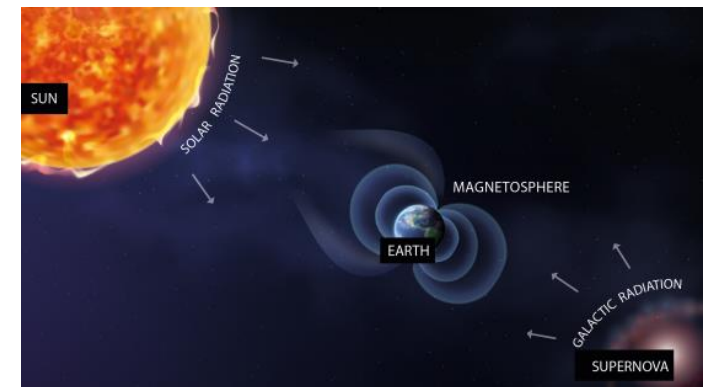


Image from: <https://www.iaea.org/newscenter/news/cosmic-radiation-why-we-should-not-be-worried>

Modelling equivalent doses and particle fluxes inside the ISS with OLTARIS

- On-Line Tool for the Assessment of Radiation in Space (OLTARIS) is an internet-based tool to assess the risk of radiation on astronauts or equipment in ISS or in spaceship
- Runs on NASA's High Charge(Z) and Energy TRaNsport (HZETRN) deterministic code



User Name
Password

OLTARIS
On-Line Tool for
the Assessment of
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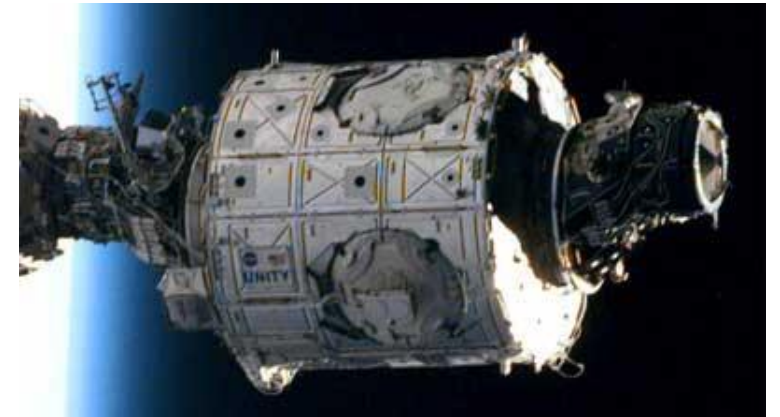
[+ Documentation and Links](#)



Thickness distributions

Built-in ISS components:

- US LAB
- Node 1
- Cupola
- Service module



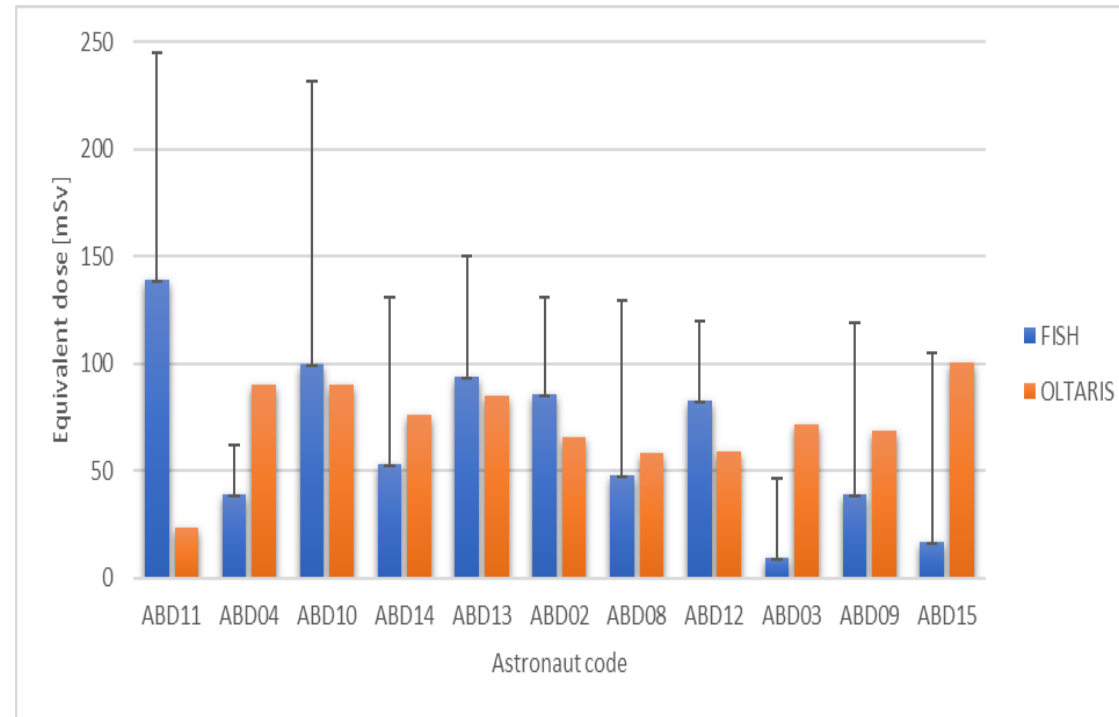
Simulation parameters

Parameter	Feature Used
Environment	Circular Earth Orbit
Altitude	400 km
Inclination	51.6 degrees
GCR model	Badhwar-O'Neil 2020
Trapped particle model	AP8
Dose equivalent quality factor	ICRP60

Comparison of OLTARIS and HC biodosimetry dose estimates

- Dose estimates from biodosimetry are usually higher than OLTARIS
 - Possibly due to using low LET calibration curves to model predominately high LET radiation.

- With permission and consent from CSA, ESA, and NASA medical boards and individuals, HC would like to benchmark these results with physical dosimetry

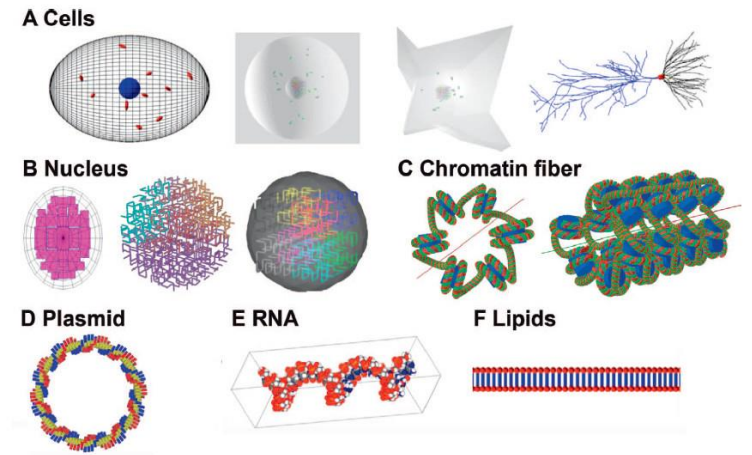


Code	Dose Rates OLTARIS [mSv/day]
ABD11	0.47
ABD04, ABD10	0.48
ABD14	0.47
ABD13	0.44
ABD02	0.38
ABD08	0.35
ABD12	0.35
ABD03	0.36
ABD09	0.37
ABD15	0.49

Modelling biodosimetry with Monte Carlo programs

Modelling biodosimetry: TOPAS-nBio

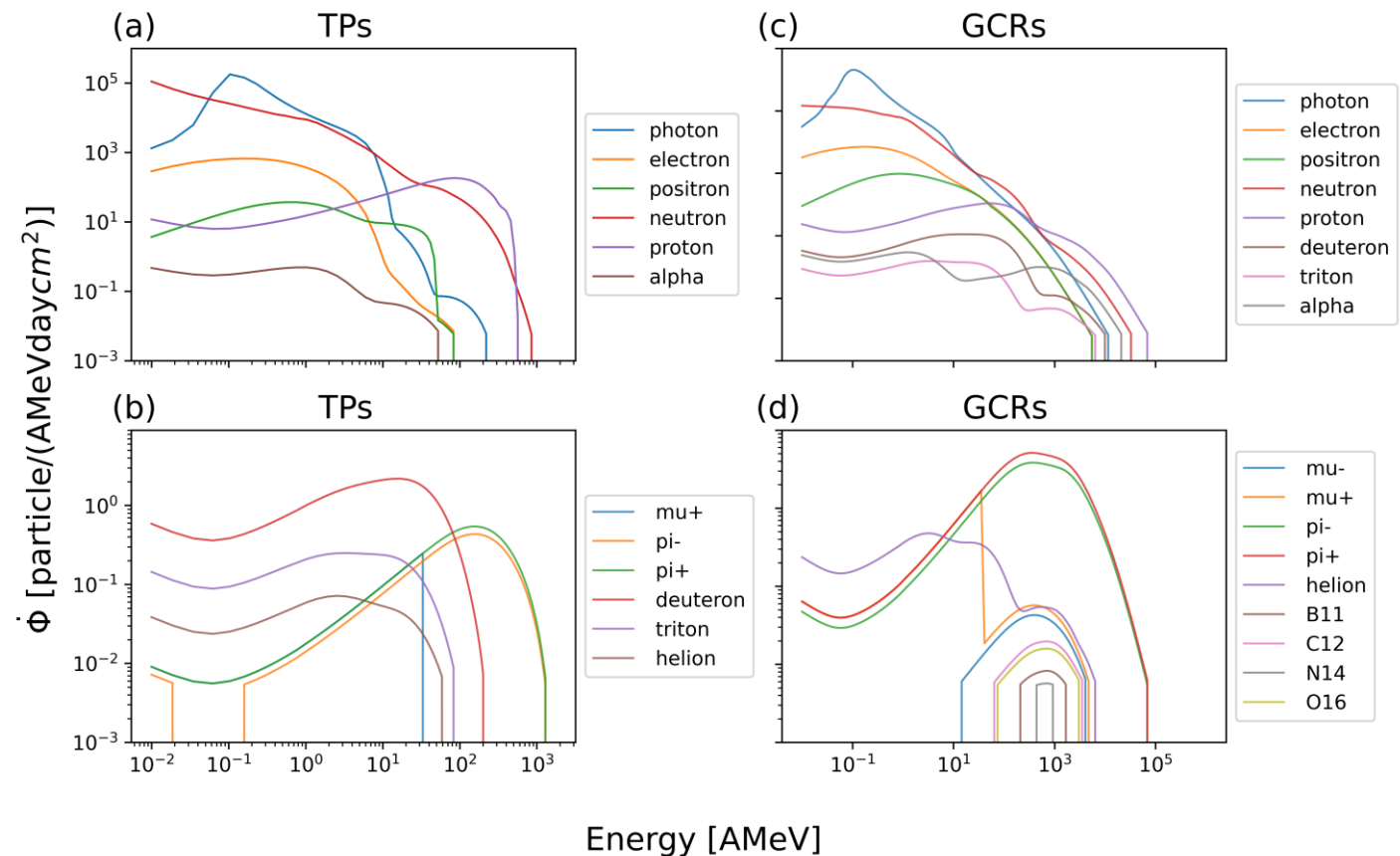
- TOPAS-nBio is Monte Carlo tool package that is built off and extends Geant4-DNA
 - Focusses on biological responses to radiation damage down to the DNA level
 - Can implement different cell types and geometries
 - Capable of outputting scored DNA damage in the standard DNA damage format



Schumann et al. RADIATION RESEARCH 191, 125–138 (2019)

Modelling biodosimetry: mixed beam calibration curves

- Produced particle fluxes as function of energy for TPs and GCRs for 2015 mission in Cupola component
 - Tune these particle fluxes and use as source inputs (e.g. to irradiate a tissue equivalent phantom) in Monte Carlo programs to produce mixed beam calibration curves
- With these simulated mixed beam calibration curves, we will determine a correction factor
- Correction factor will help improve the accuracy of our current assay



Conclusions

- ✓ Discussed and presented current state of Health Canada's space biodosimetry programme
- ✓ Simulated astronaut missions from the programme with NASA's OLTARIS to obtain equivalent dose estimates and particle fluxes
- ✓ Equivalent dose estimates from OLTARIS are generally lower than those obtained by HC's biodosimetry

Next steps

- ❑ Perform these mission simulations with ESA's SPENVIS program to validate the particle flux results from OLTARIS
- ❑ Work is underway to build mixed radiation beams with Monte Carlo programs (TOPAS-nBio or other ...) to obtain mixed radiation calibration curves
- ❑ Benchmark mixed radiation calibration curves against experimental ones

Acknowledgements

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