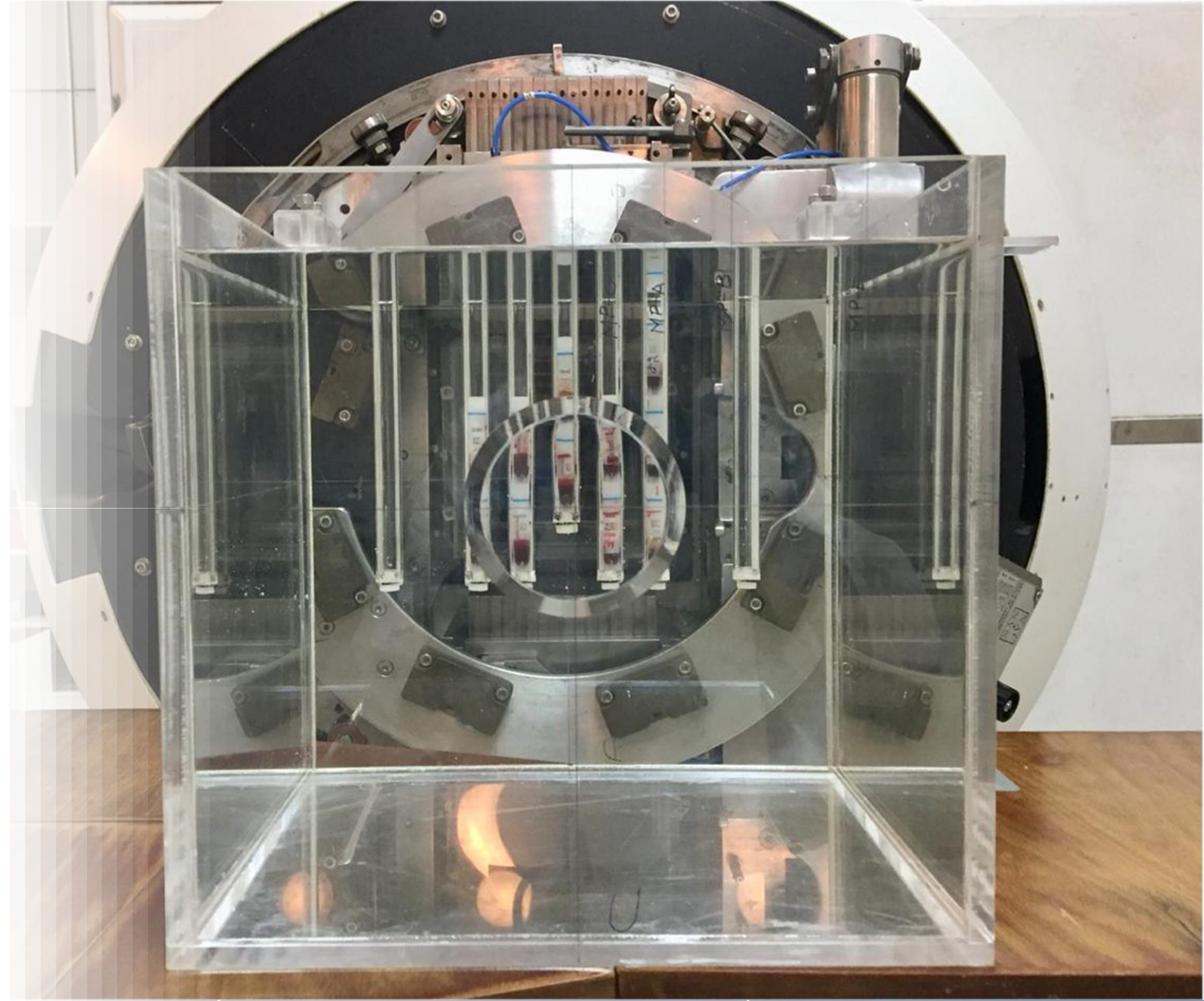


RADIATION BIOPHYSICS

A division of the Separated
Sector Cyclotron Laboratory
NRF iThemba LABS

Dr Randall Fisher
rg.fisher@ilabs.nrf.ac.za



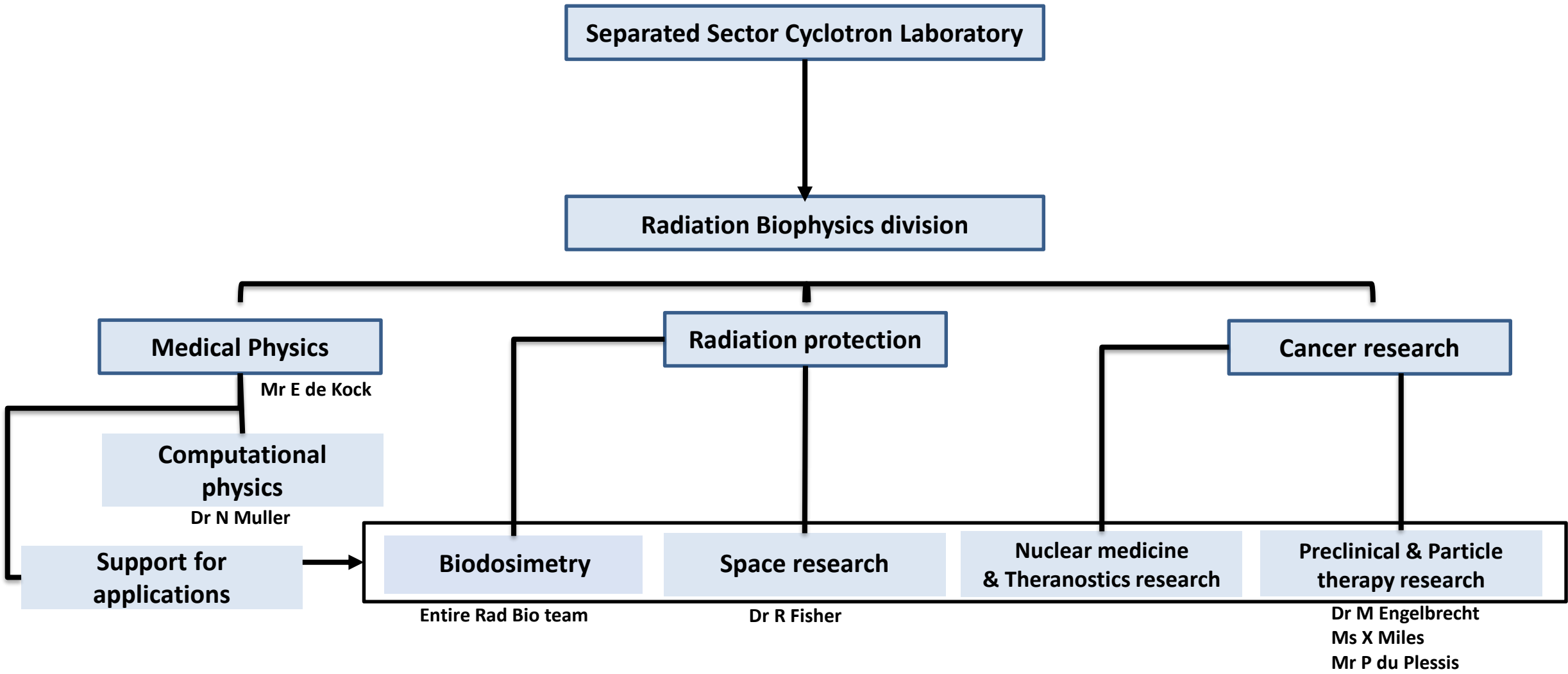
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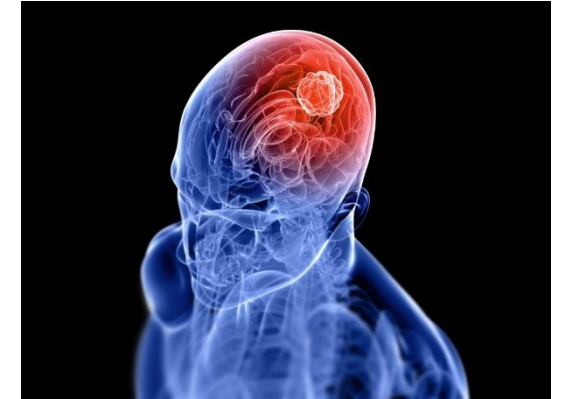
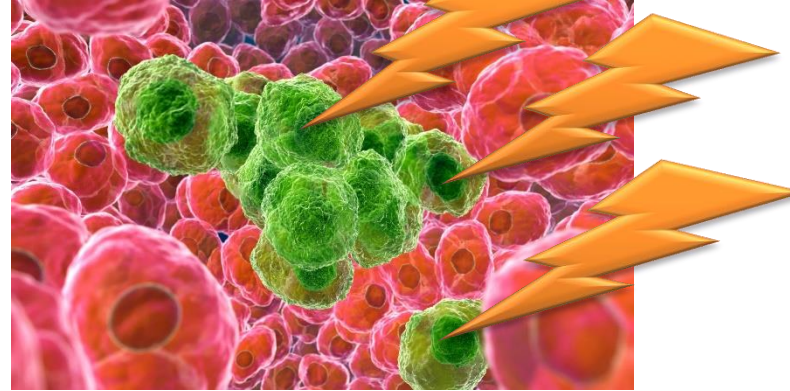
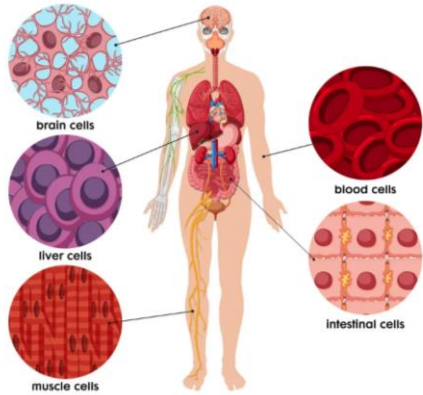
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LABS
Laboratory for Accelerator
Based Sciences



What is Radiobiology?

Cells of The Human Body



- Our projects investigate:
 - Using drugs to improve the recovery of normal cells or improve the killing of cancer cells
 - Different radiation types, what they do to cells and how they can be used to improve cancer therapy
 - The way that different radiation environments affect the human body and genetic cancer
 - New ways to identify and treat cancer with radioactive drugs
 - Genetic mechanisms of cancer resistance



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RADIATION BIOPHYSICS

Current projects



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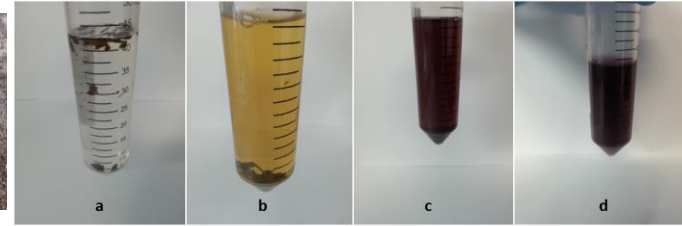
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LABS

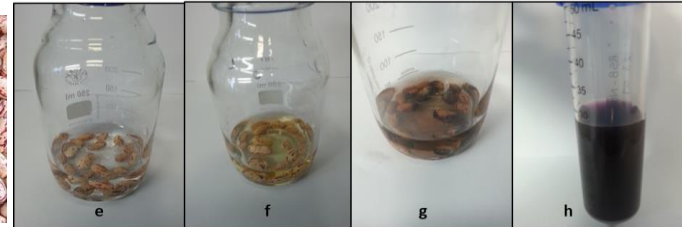
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Preclinical - Radiosensitisers

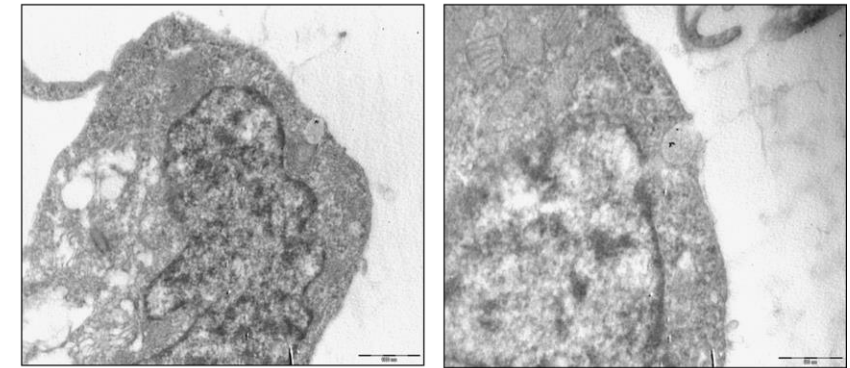
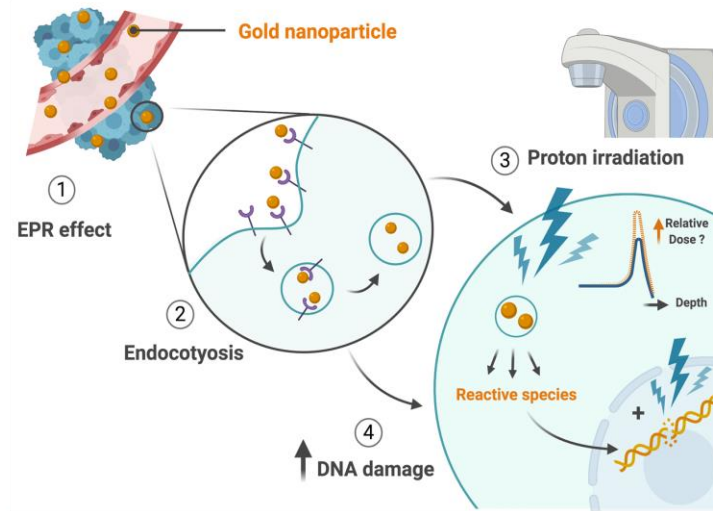
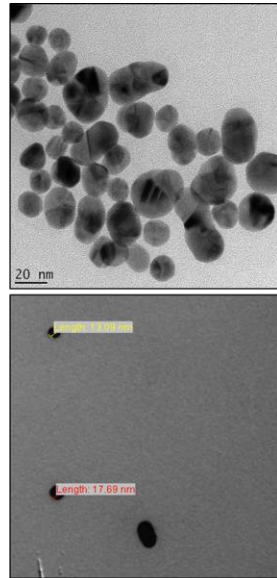
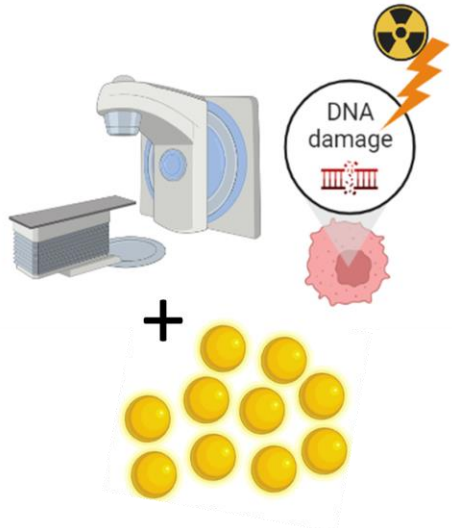
- Gold/Green NPs
- Radiosensitising drugs – AMG232, CUDC-101, Vorinostat (SAHA), Endostatin



Cinnamon gold nanoparticles (Cin-AuNPs)



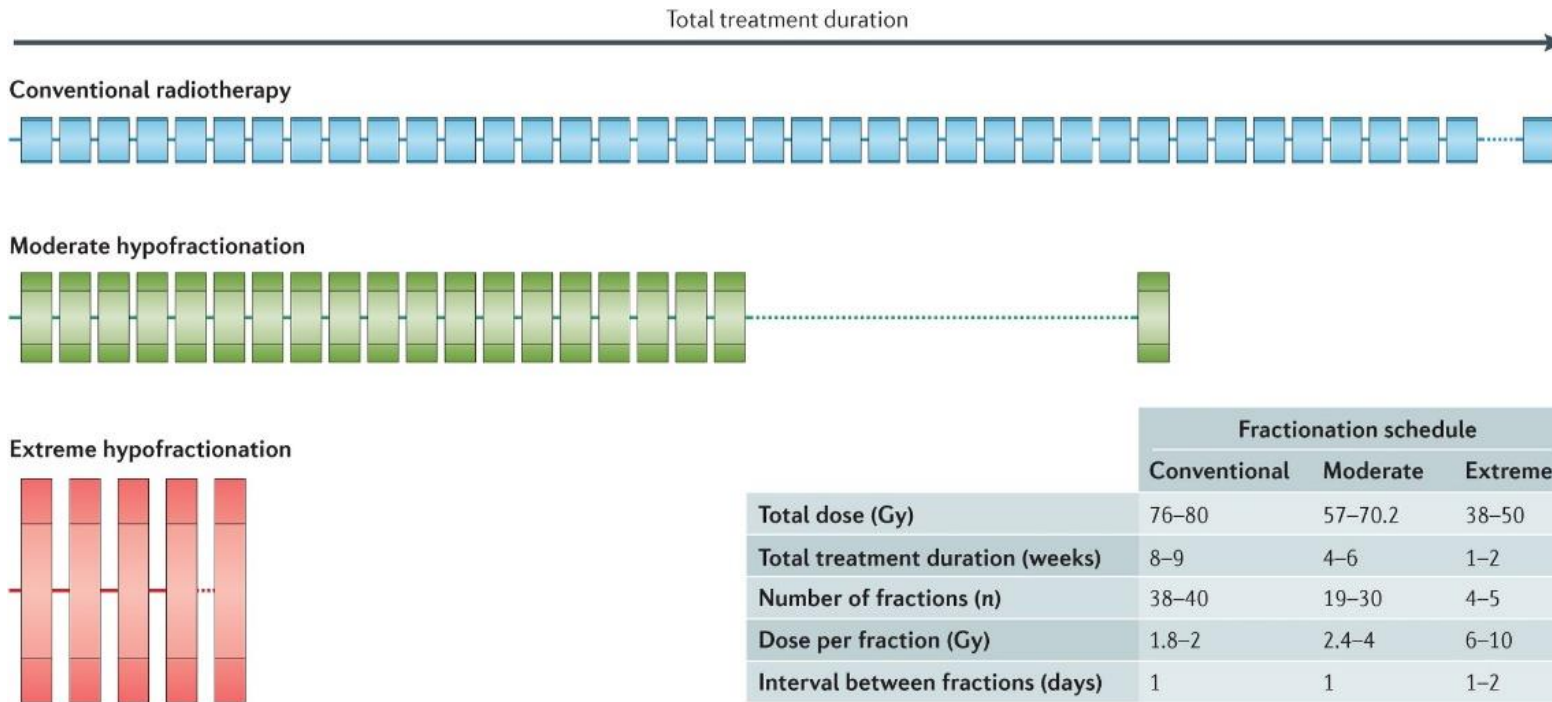
Red speckled beans gold nanoparticles (RSB-AuNPs)



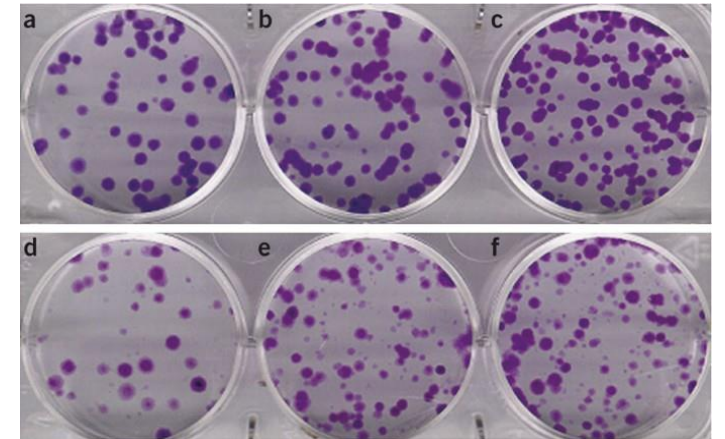
More effective therapy

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Preclinical – Hypofractionation

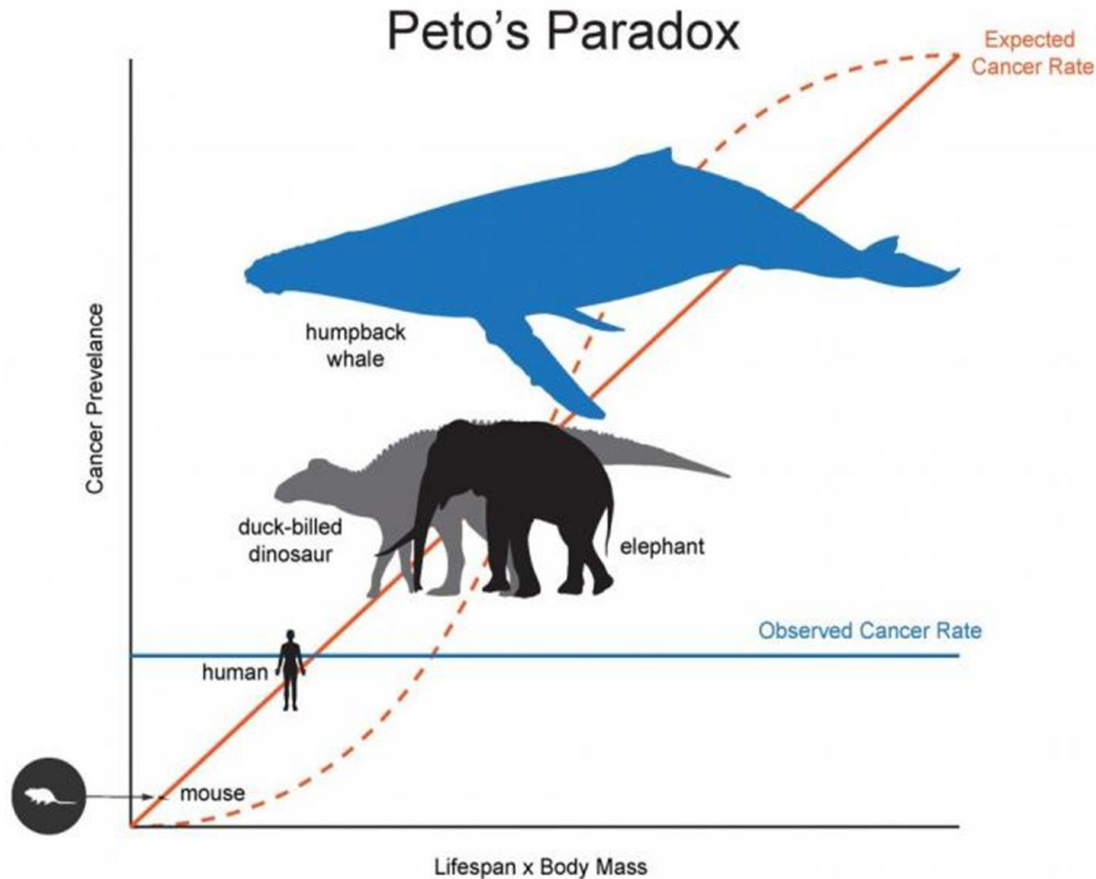


- Loss to follow-up in the clinic breeds resistance
- Hypofractionation should achieve a biological effective dose



Nature Reviews | Urology

Preclinical - TUmour Suppression and Subdual of Cancer (TUSSC) in elephants: An *in vitro* study to shed light on Peto's Paradox



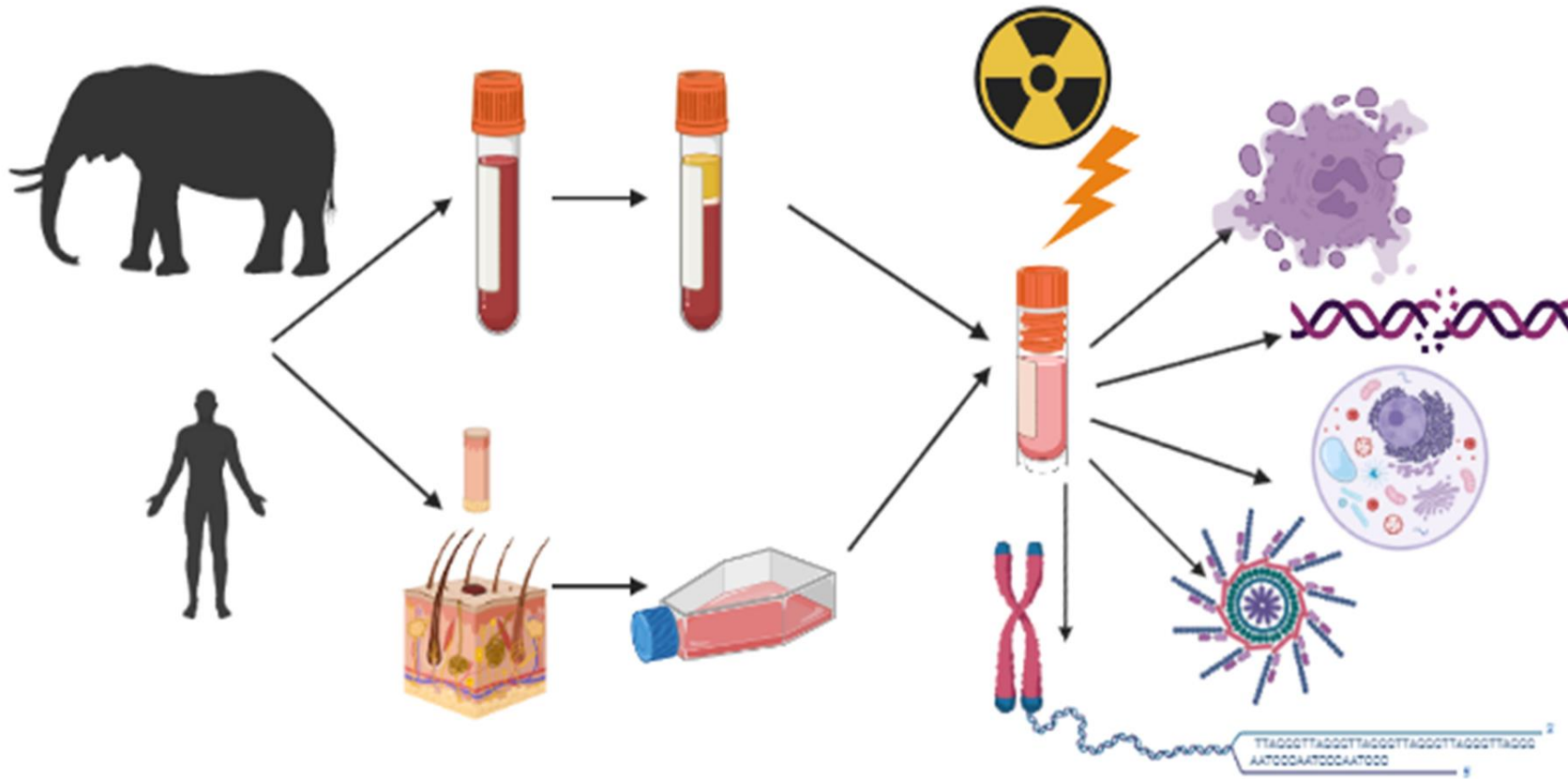
70 years	Average Life Span	>60 years
65 kg	Average weight	4800 kg
40 billion	Number of cells	40 trillion
25%	Cancer mortality	<5%
1	Copies of p53 gene	20

PROJECT TUSSC

TUMOUR SUPPRESSION AND
SUBDUAL OF CANCER

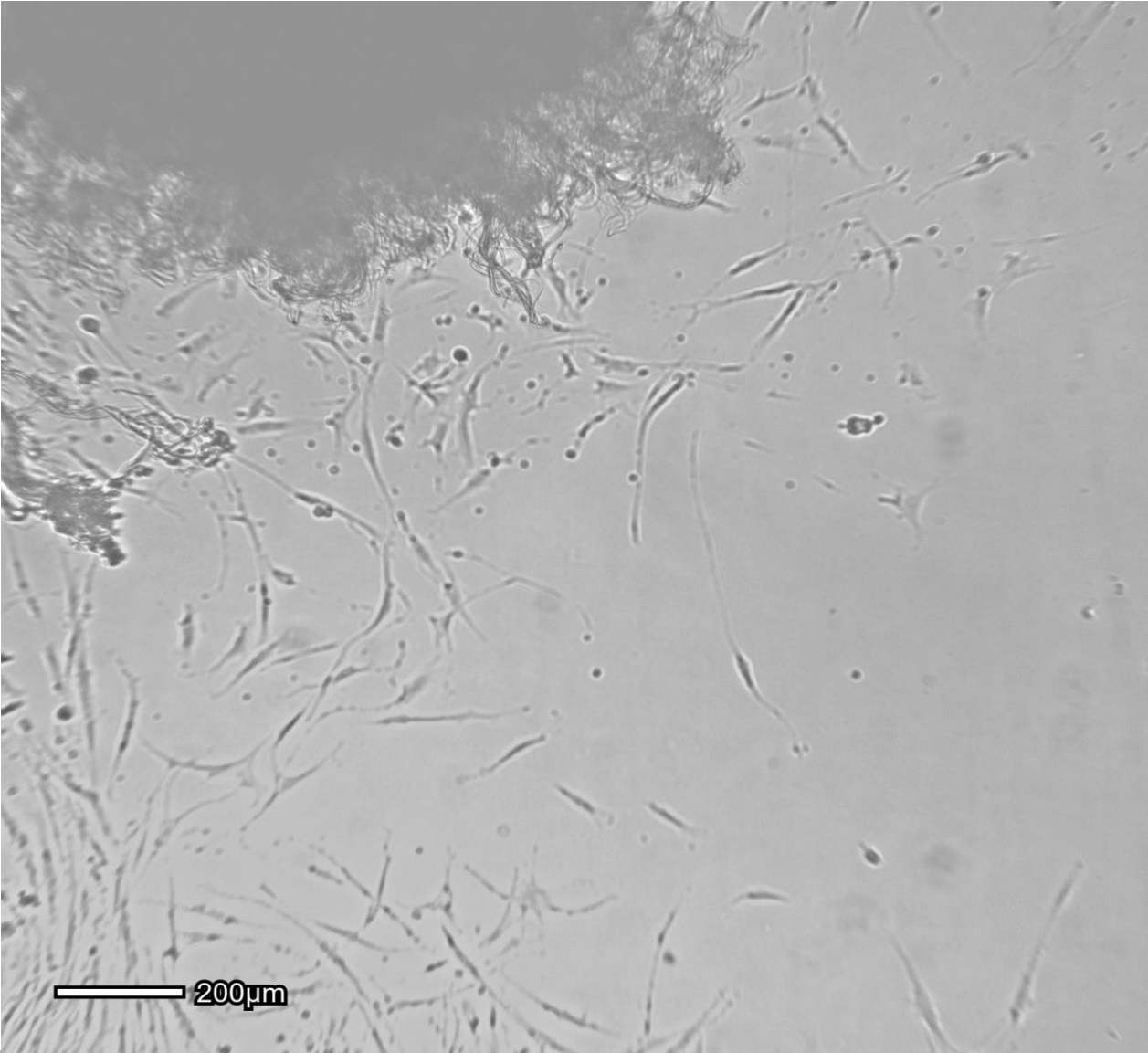
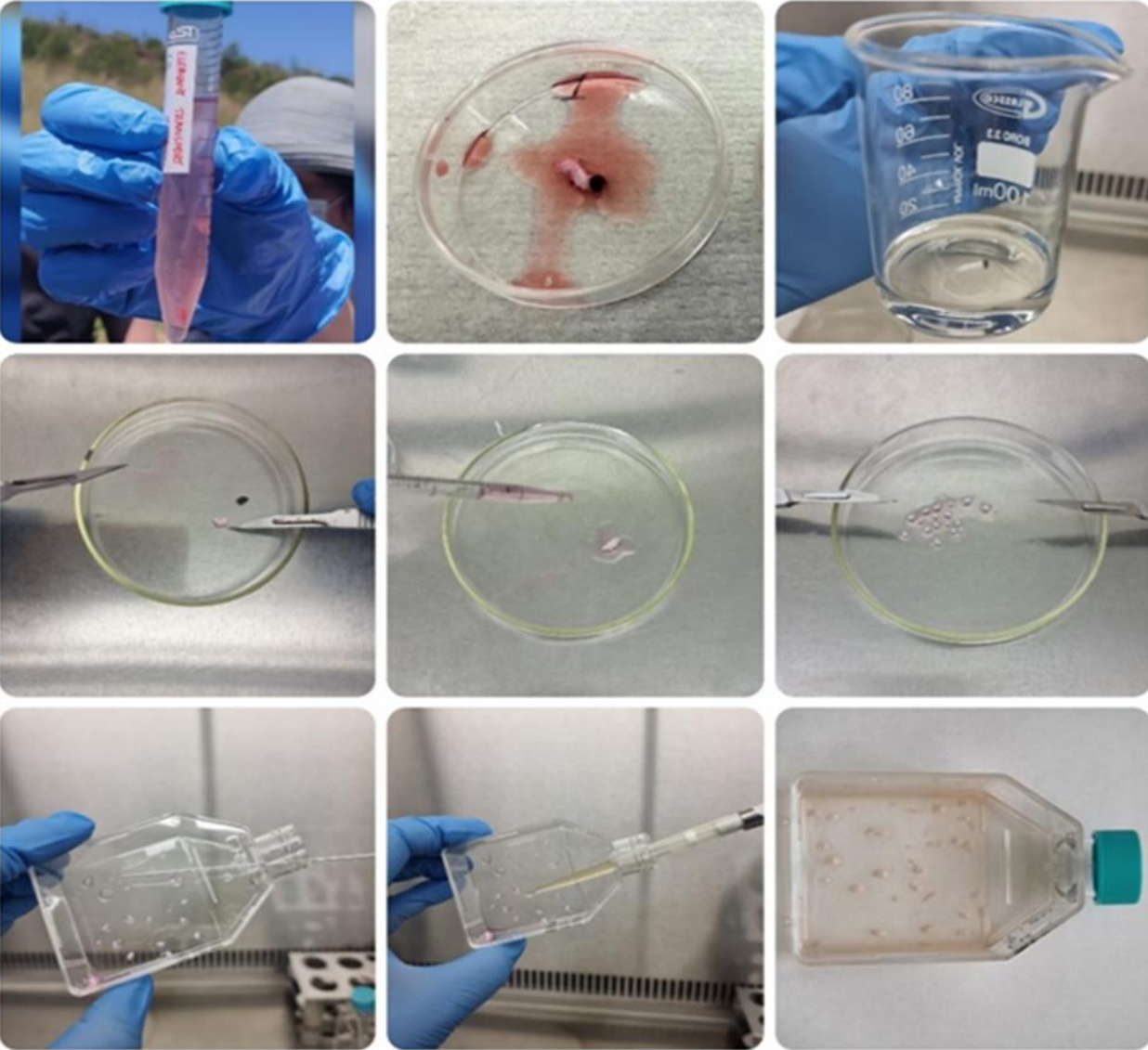


Preclinical - TUSSC Project



- Apoptosis
- DNA repair
- Metabolism status in the cancer suppression of elephants
- Inflammasome pathway of the immune system
- Role of telomere length and telomere shortening rate in Peto's paradox

Preclinical - TUSSC Project



Biodosimetry and Radioprotection – Spaceflight Radiobiology

- How to minimise the health risks induced by radiation exposure?

Space Radiation = Galactic Cosmic Rays and Solar Energetic Particles.

Extreme conditions: cosmic radiation, prolonged weightlessness and social isolation.

Causes stress, affect our eyesight, make bones more brittle.

Increase the risk of cancer and heart diseases.

Little known on the health effect of these space stress factors with psychological stress.

A space trip to Mars takes about two years.

Zero gravity



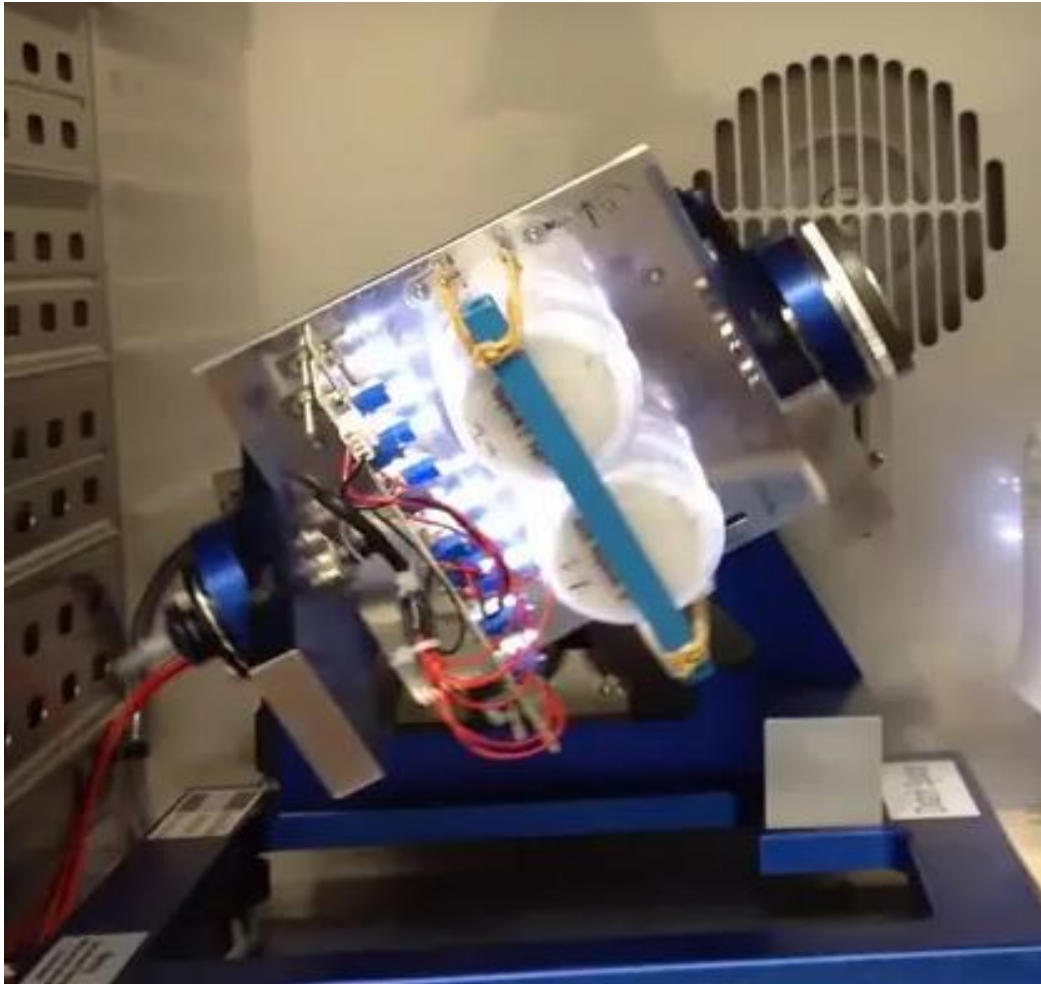
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Biodosimetry and Radioprotection – Spaceflight Radiobiology



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Future projects



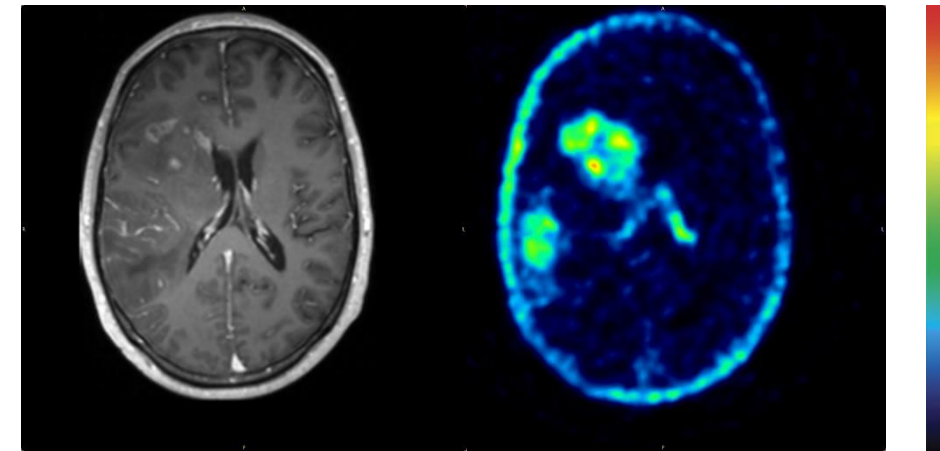
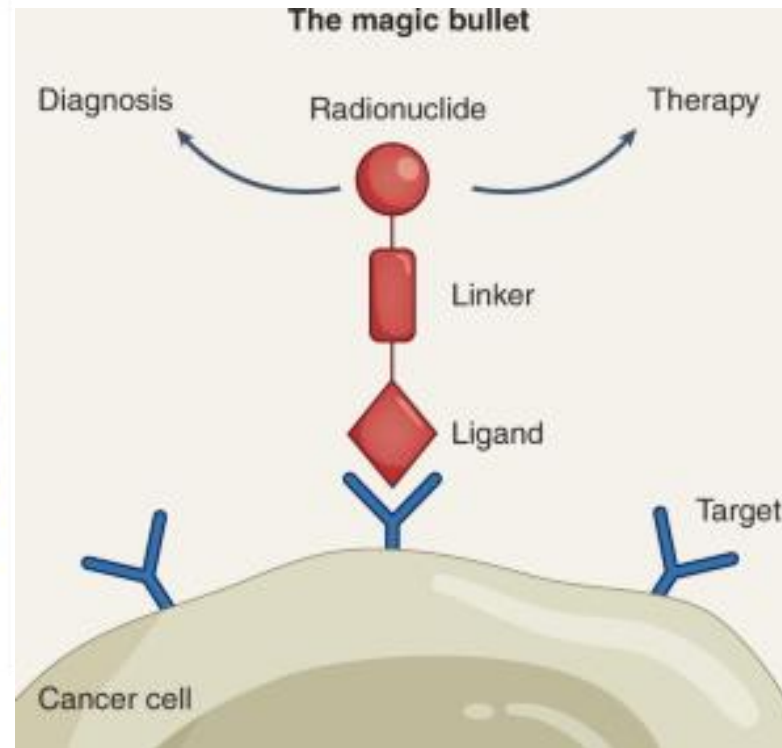
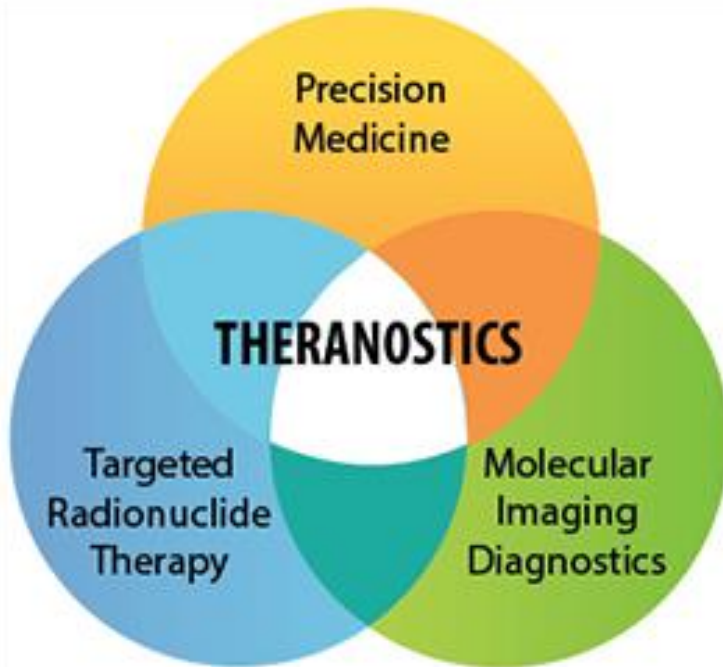
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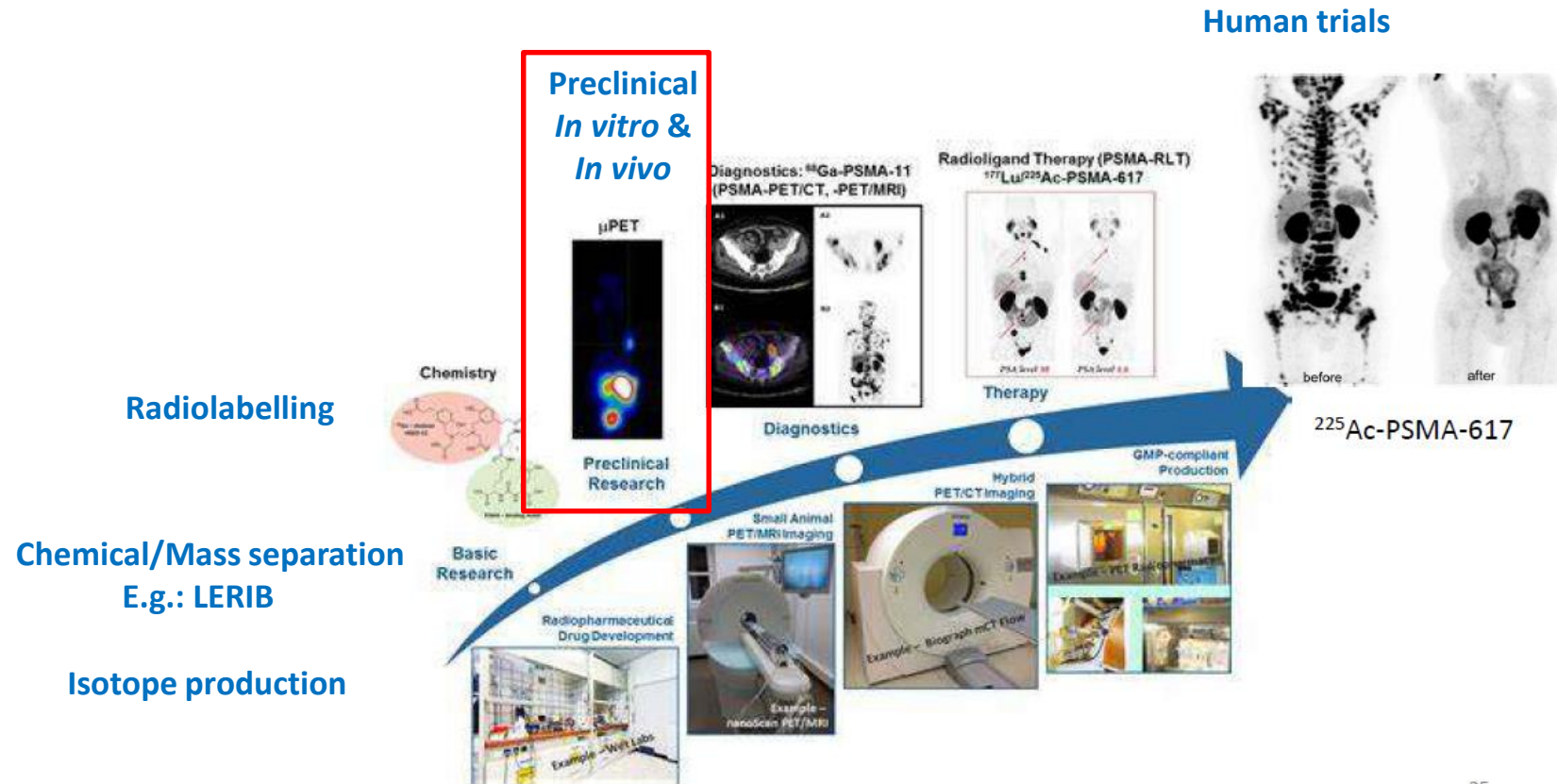


Theranostics

= therapeutics + diagnostics



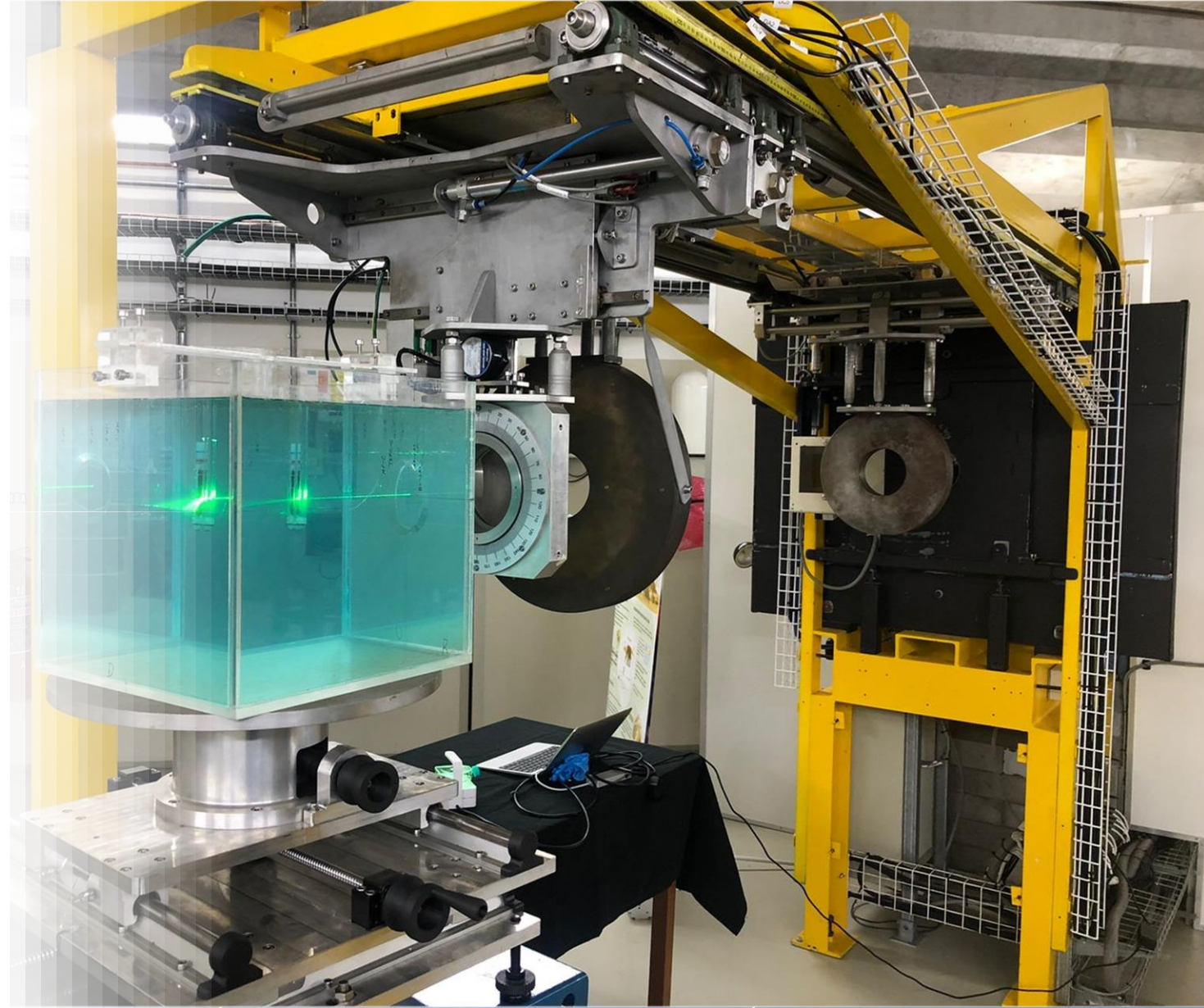
Pipeline for Theranostics research



25

RADIATION BIOPHYSICS

Infrastructure



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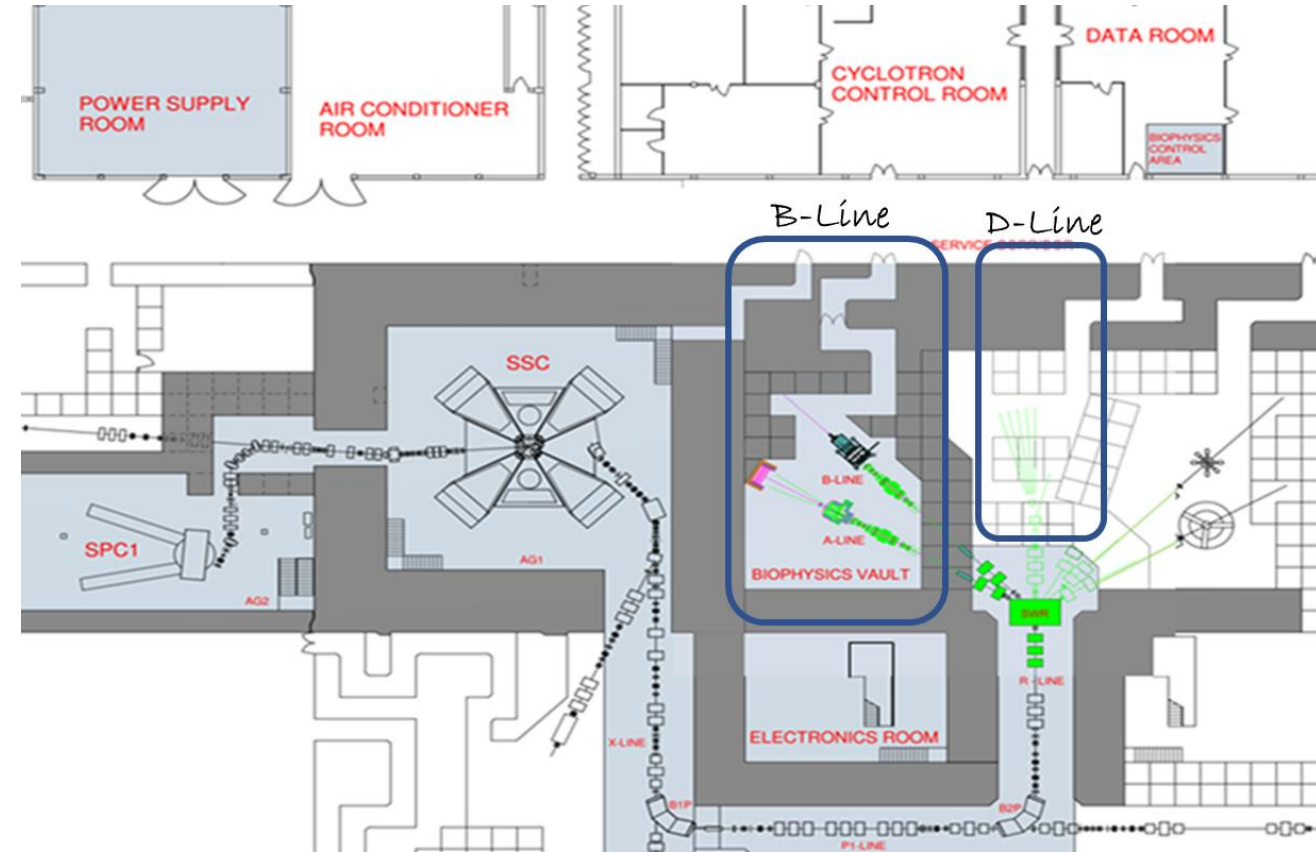
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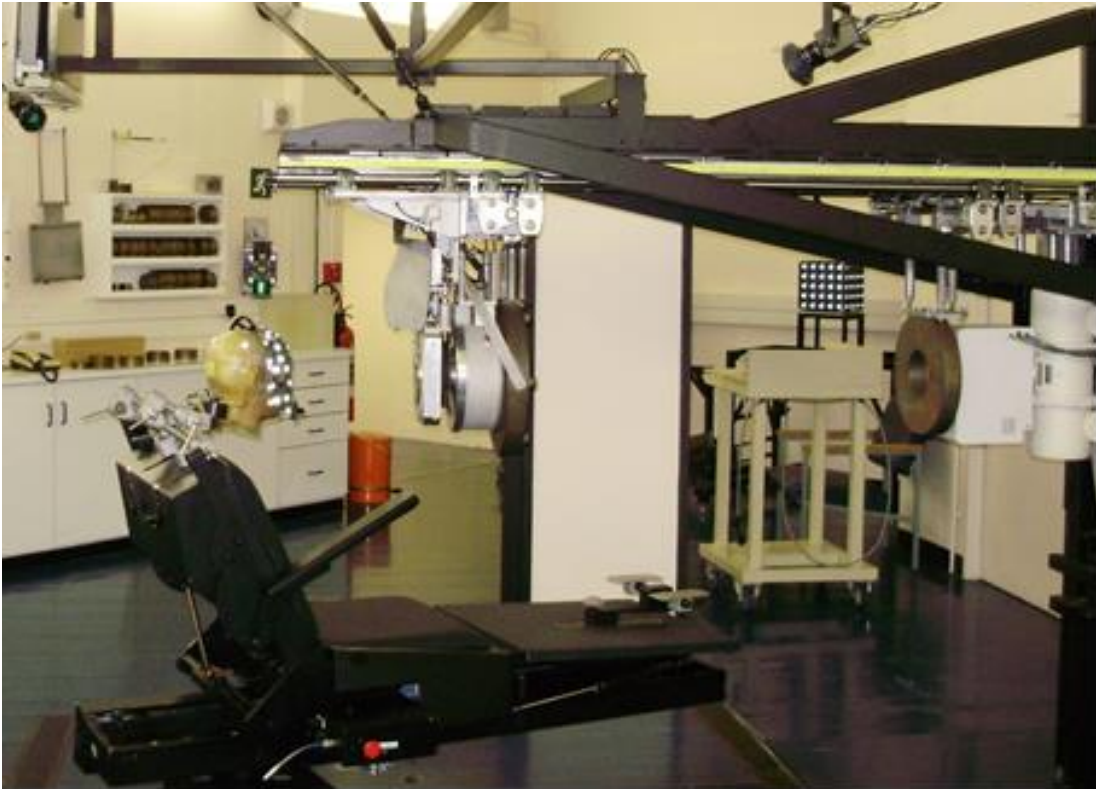
Infrastructure: Irradiation facilities

- Horizontal 200 MeV proton therapy beam delivery system in the B-line,
- A Quasi-mono energetic neutron beams produced in the D-line
- Self-contained biological X-ray irradiator (X-Rad320, Precision X-ray)
- Cobalt-60 source (from an older Theratron 780 teletherapy unit)
- The $p(66)/\text{Be}$ isocentric neutron therapy unit is no longer available.



Infrastructure: Irradiation facilities

Horizontal 200 MeV proton therapy beam delivery system in the B-line



Infrastructure: Irradiation facilities

Horizontal 200 MeV proton therapy beam delivery system in the B-line

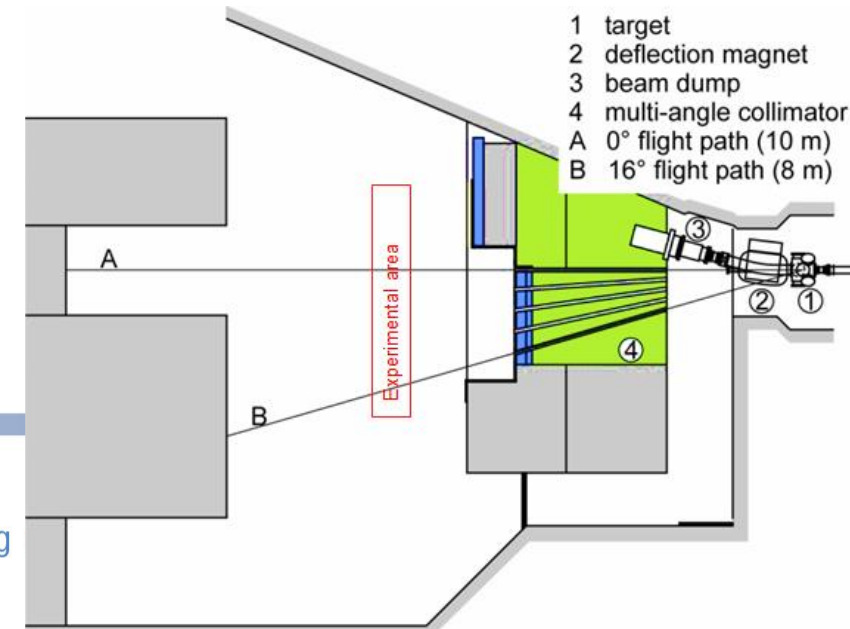
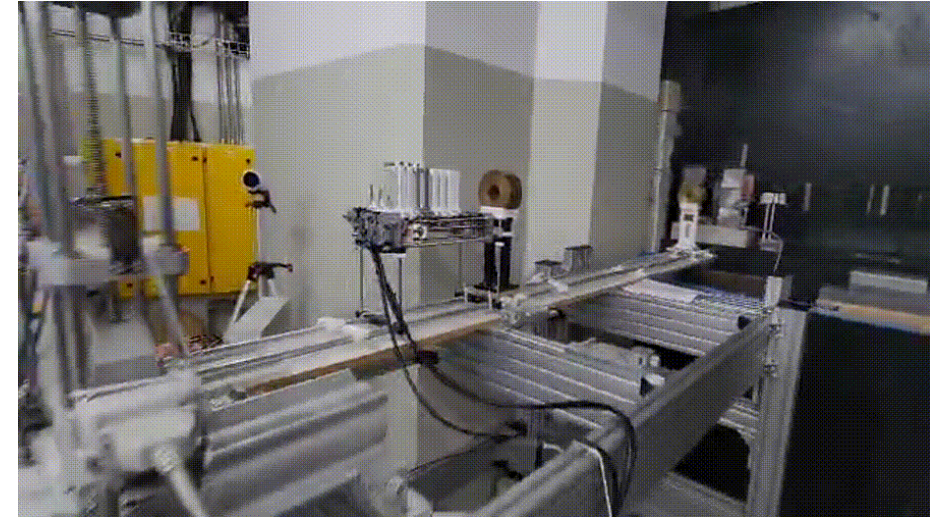
- A passive double scattered and occluding ring system is used to spread and flatten the beam.
- Graphite-wedge flattening filters.
- Maximum field of 10 cm in diameter.
- The reference beam used for absolute dose calibrations have dose rate of 3.5 Gy/min at depth of 3 cm in water.
- Incident proton beam current of 20 nA – 100 nA.
- FLASH dose rates (>40 Gy/sec)



Infrastructure: Irradiation facilities

Quasi-mono energetic neutron beams produced in the D-line

- Medium- to high-energy neutron metrology facility
 - Quasi-monoenergetic neutron beams with energies 30 – 200 MeV, using (p,n) reaction on thin Li and Be targets
- Collaboration with UCT MeASURe group
 - 5x5 cm field at 0°, 4°, 8°, 12° and 16°
 - Dose rate **0.04 Gy/hour (1.2μA) to max 0.7 Gy/hour**
 - These beams are useful for the characterisation of dosimetry detectors (neutron energies up to 200 MeV). Future developments are planned to cater for *in vitro* radiobiology.



Infrastructure: Irradiation facilities

Self-contained biological X-ray irradiator (X-Rad320, Precision X-ray)

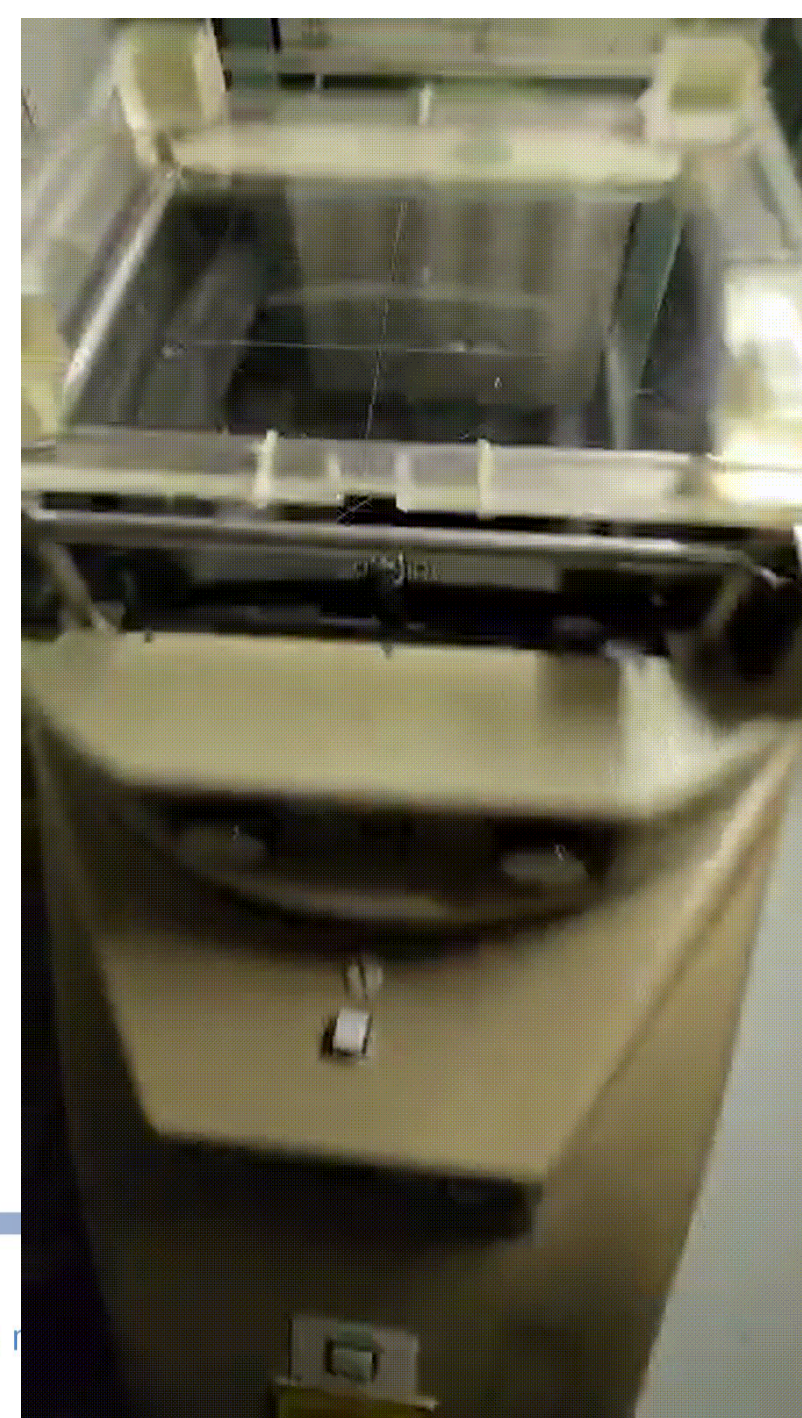
- In close proximity to the cell culture facilities.
- 250 kV commissioned for four cell culture receptacles (TG-69 protocol)
 - H-series filtering (3.5mm Be, 4mm Al, 3mm Cu)
- 5 energy ranges: 100 kV to 300 kV
- Field size set as 19 cm × 19 cm at a SSD of 50 cm.
- The average dose rate for the T25 flask (for the standard setup in the 250 kVp beam) is 0.685 Gy/min for a tube current of 16 mA. 5 energy ranges (100 to 300 kV)



Infrastructure: Irradiation facilities

Cobalt-60 source (from an older Theratron 780 teletherapy unit):

- Reference beam for *in vitro* studies
- Detector cross calibration
- ~1.25MeV
- SSD – 600 mm
- Dose rate of 0.22 Gy/min for a 27 × 27 cm² field size.



Infrastructure: Irradiation facilities

- Sterile cell culturing facilities.
- Microscopy infrastructure:
 - Two Metafer automatic scanning and imaging platforms (Metasystems)
 - Zeiss Axioscope fluorescent microscope
 - Live cell imaging system (Lonza Cytosmart)
- Flow cytometry (BD Accuri C6 flow cytometer)
- Fluorescent in situ hybridization (FISH) for chromosomal aberration analysis
- Radiochemical synthesis and validation of radiolabelled compounds:
 - Radiosynthesis laboratory
 - Rotary evaporator
 - Scintillation counter
 - High-performance liquid chromatographer
- UV-VIS spectrophotometer



Thank you for your attention!

BOTLIERSKOP
Private Game Reserve



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WESTERN CAPE



UNIVERSITEIT
STELLENBOSCH
UNIVERSITY



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