



Radiobiological Research in the Laboratory of Radiation Biology

Introduction to LRB

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JINR history and present



- 1959 first radiobiological experiments LNP
- 1978 Sector of Biological Research
Laboratory of LNP
- 1988 Department of Biophysics at LNP
- 1995 Department of Radiation and
Radiobiological Research of JINR
- 2005 Laboratory of Radiation biology





Laboratory of Radiation biology



- Molecular Radiobiology
- Radiation Cytogenetics
- Group of Lower Eukaryote Radiation Genetics
- Photoradiobiology
- Mathematical modeling
- Astrobiology
- Radiation Neurochemistry
- Radiation Physiology

Research fields of LRB

- Radiation genetic studies – mammalian and human cells
- Radiation research and development
- Computer molecular modeling of biophysical systems
- Radiobiological studies of the Retina and Crystalline lens, cataract induction, visual pigment studies
- Cosmic radiation research, specialized research of CNS
- Astrobiological research

Research on the Biological Effect of Heavy Charged Particles with Different Energies

Nuclotron, ^{12}C 500 MeV/amu



Phasotron, protons 170 MeV

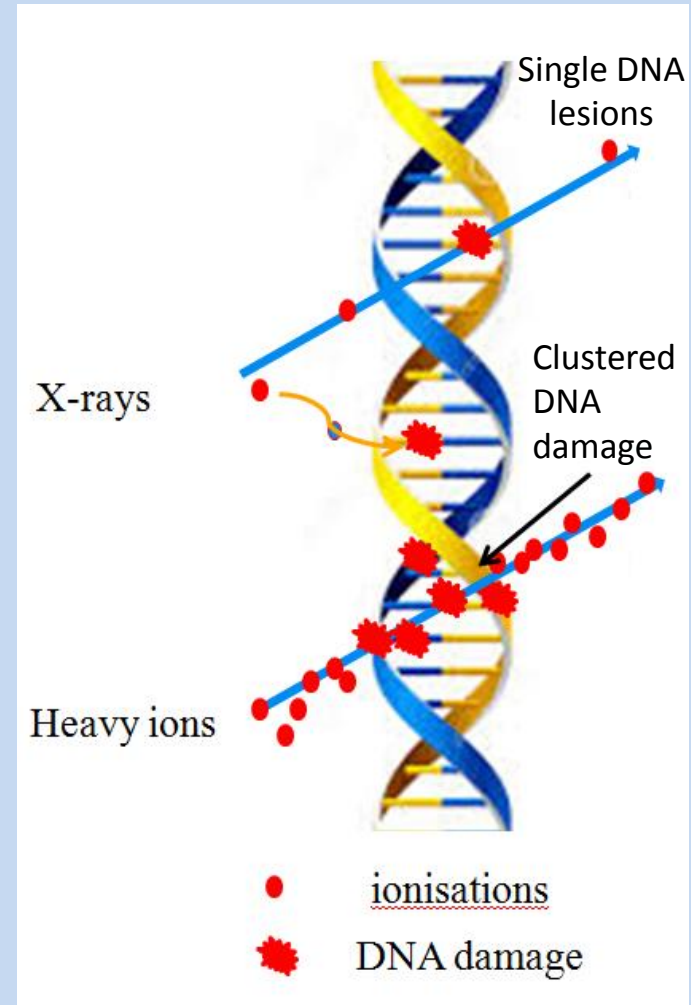
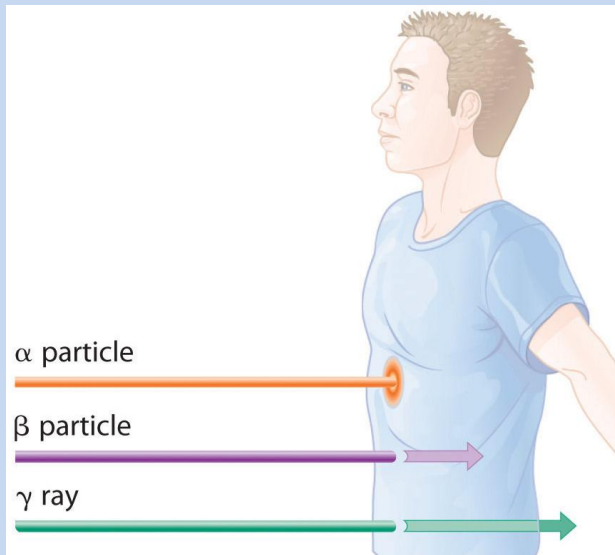
U-400M, up to 50 MeV/u Li, B, Ne ions

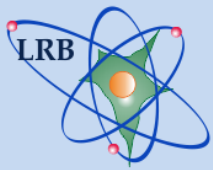




Radiation: sources and biological effects

- Electromagnetic radiation, plane travels, medical radiation, cosmic radiation
- Cell damage – free radicals formation – DNA damage – cell death/ mutations





Effects of radiation

- Early Effects:

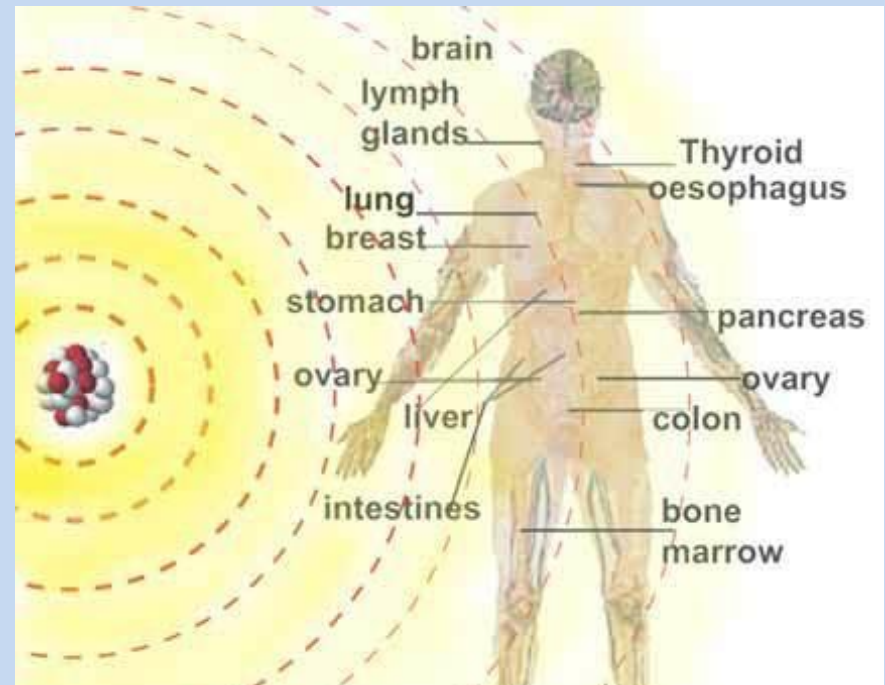
- radiation response occurs within minutes/days after exposure
- acute radiation syndrome, local tissue damage

- Late Effects:

- radiation response observable after months or years
- local tissue damage, leukemia, cataracts, cancers, CNS damage, life span shortening, sterility

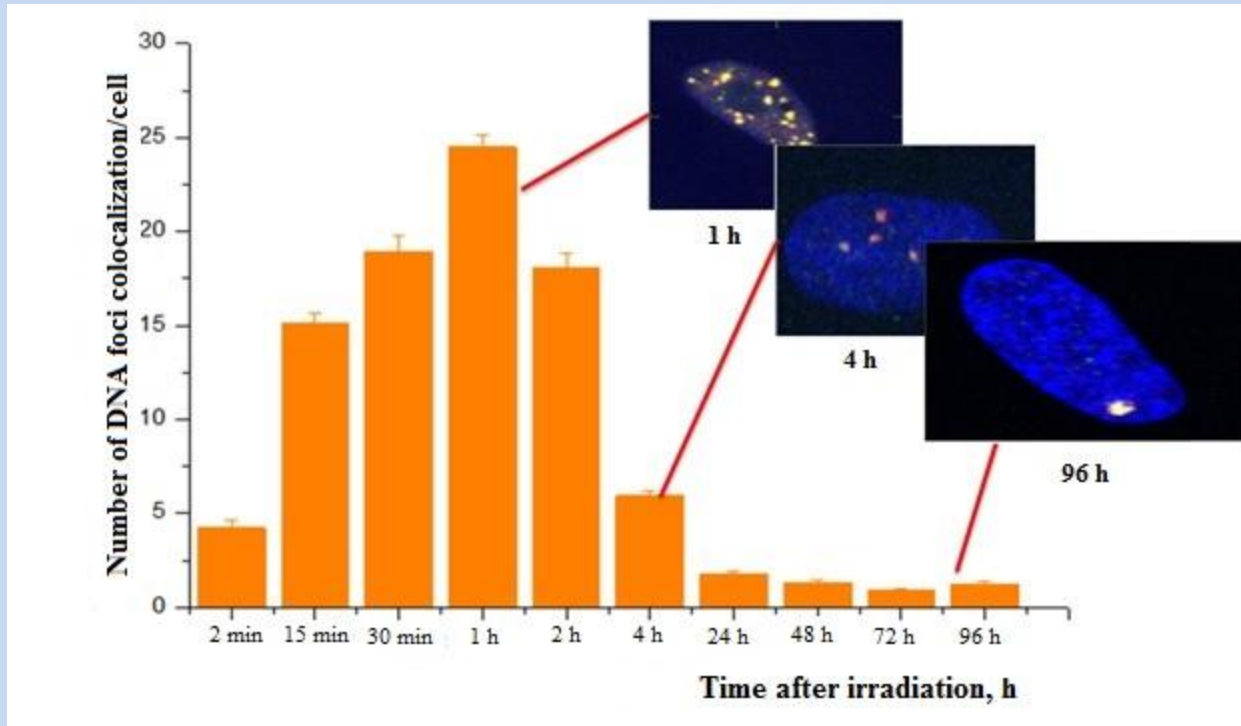
- Genetic Effects:

- teratogenesis, mutagenesis

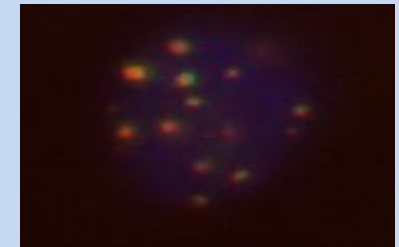


Radiation immunocytochemistry

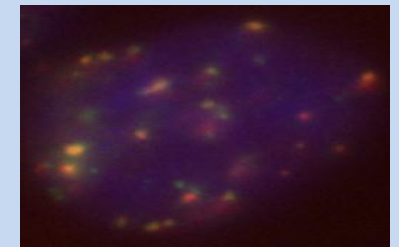
γ -rays



^{20}Ne ions, 50 MeV/nucleon



30 min



6 h

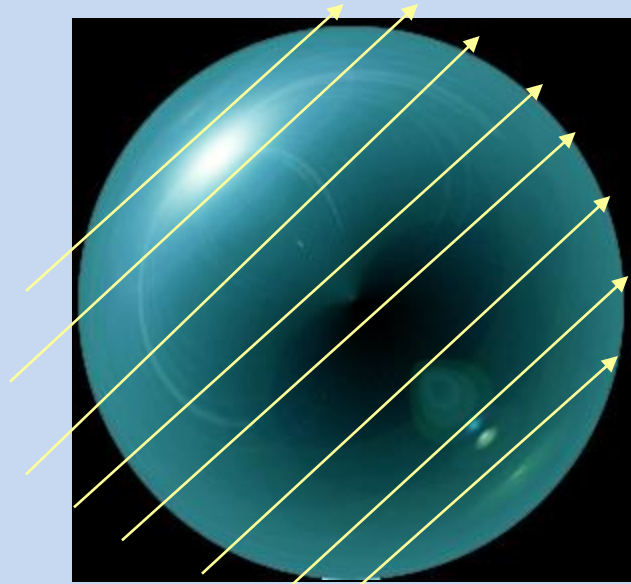


48 h

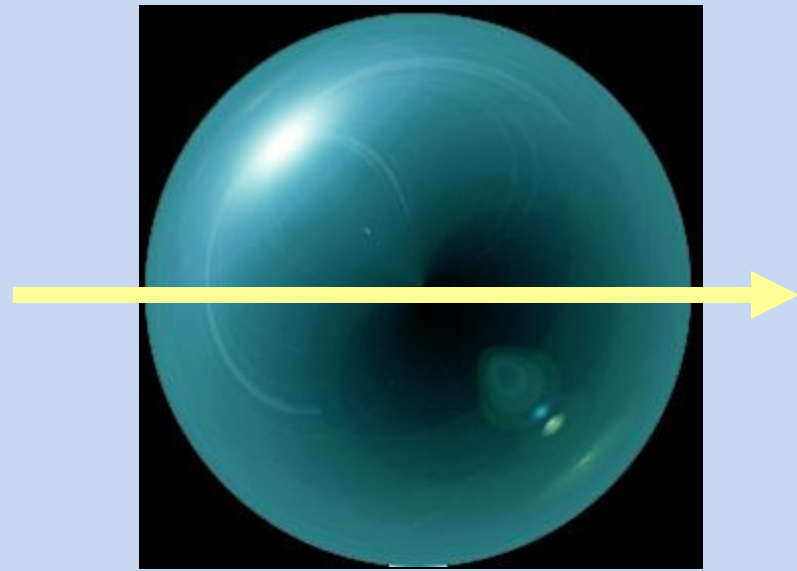
Double strand break induction (53BP1 and γH2AY foci) in human skin fibroblasts after irradiation with accelerated ^{20}Ne ions

The dose distribution of radiation in matter

1 unit of the dose

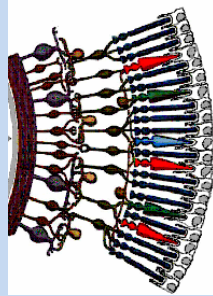
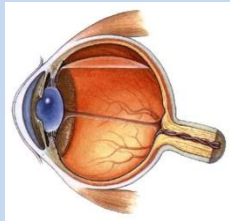


X-rays

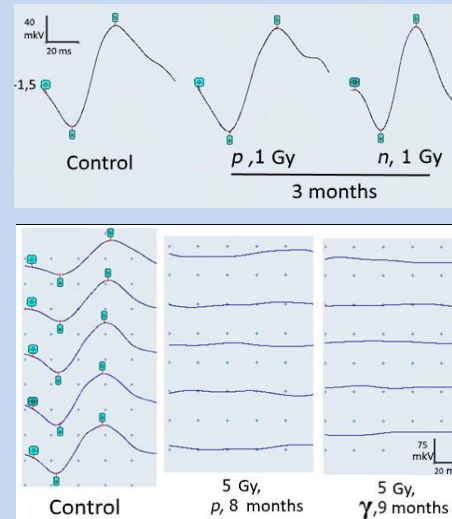
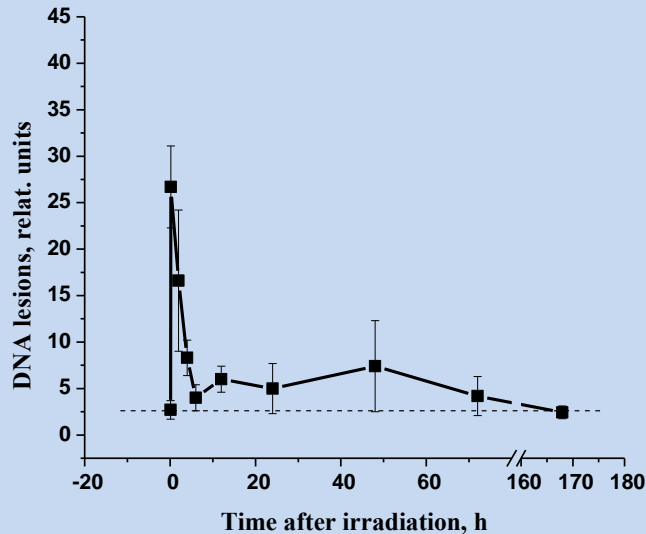


Fe ion

Damage of eye structures by accelerated charged particles

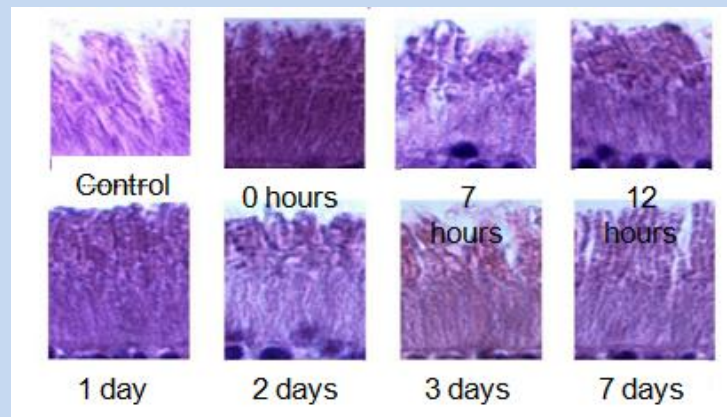


DNA lesions in retina cells after proton (170 MeV) irradiation



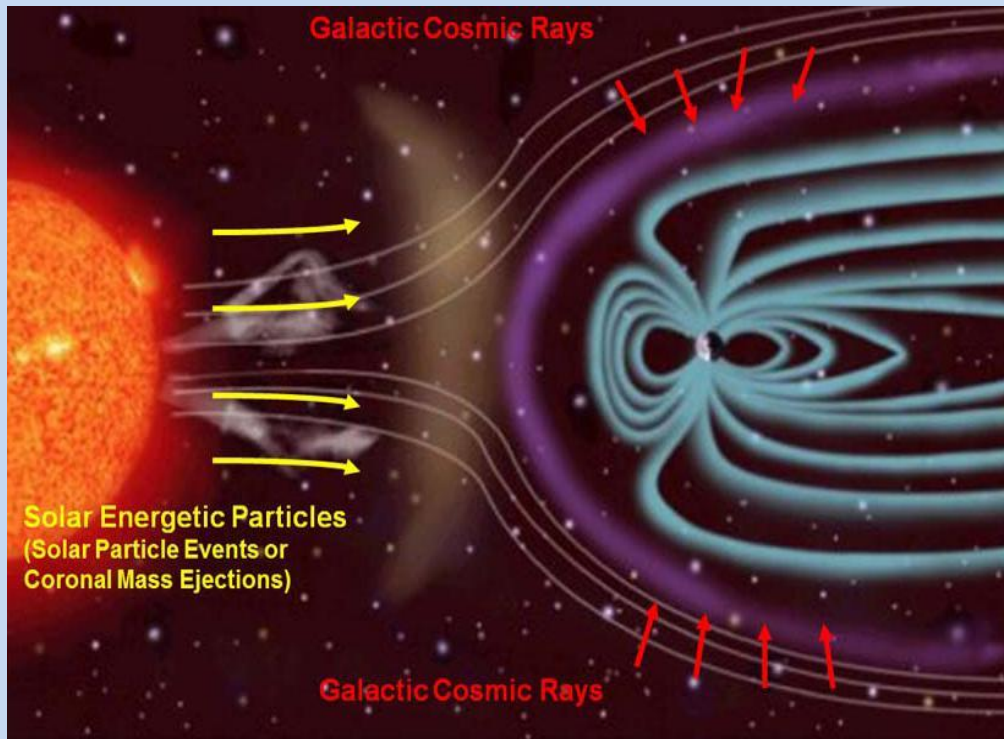
Functional activity of the eye retina of mice after exposure to low doses of ionizing radiation

Dynamics of **morphological changes** in the retina after proton irradiation

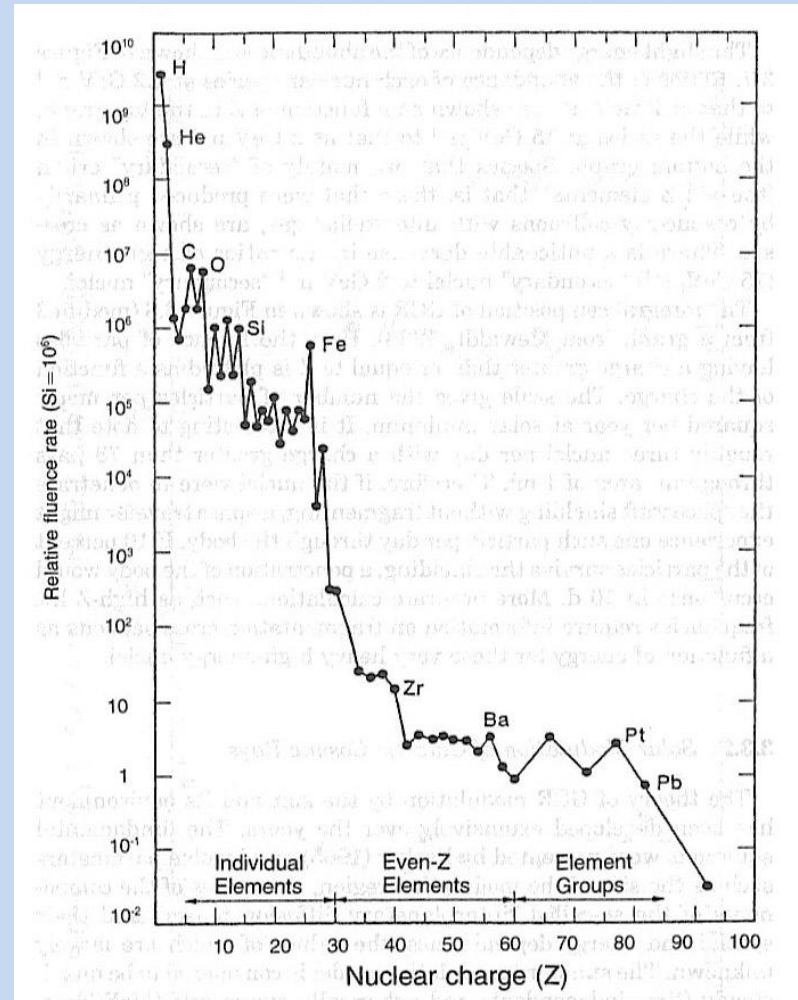


Cosmic radiation

- Solar energetic particles, galactic cosmic rays (protons-92%, nucleus of helium atoms-7%, heavy nuclei), stardust (exploding stars)
- Secondary neutrons and charged particles are the major sources of radiation exposure in an interplanetary spacecraft



Composition of Galactic Cosmic Radiation

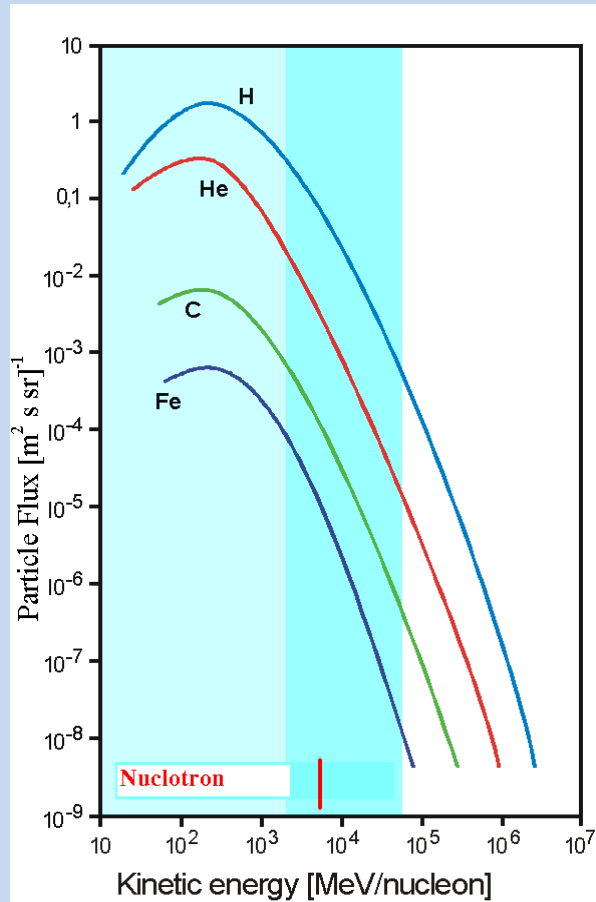




LRB Cosmic radiation research

The radiation barrier for manned missions into Deep Space and modeling of biological action of space radiation at accelerators of heavy ions

The energy spectrum of GCR



The integral flux of GCR particles of carbon and iron groups equals to 10⁵ part/cm² per year

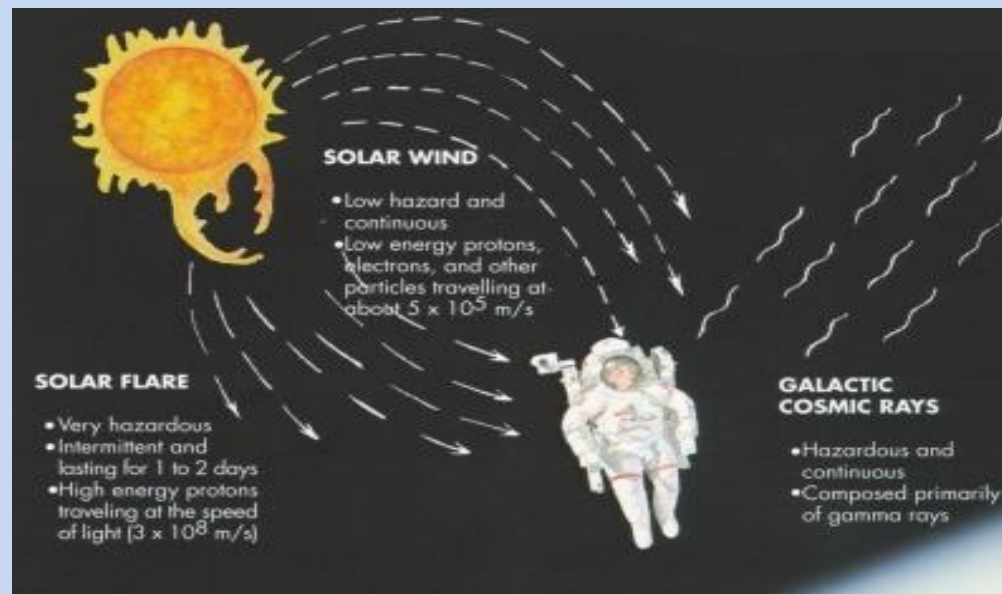
Particle flux density interplanetary space $z \geq 20$
160 per day per cm²



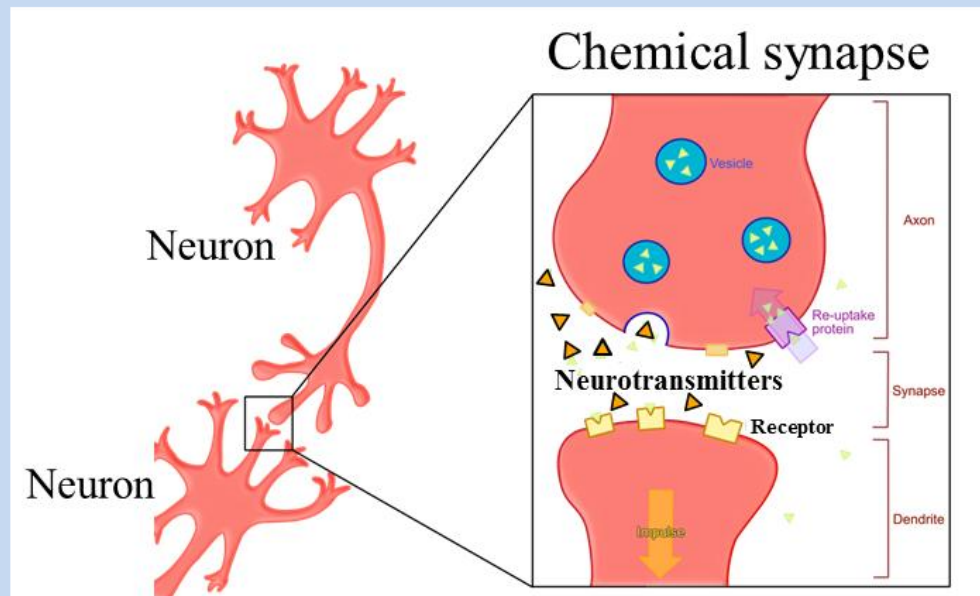
Central nervous system under cosmic radiation

During the Martian mission:

- from 2 to 3% of nerve cells will be crossed by at least one iron ion;
- from 8 to 46% of nerve cells will be crossed by at least one particle with $Z \geq 15$;
- each nucleus of the cell within 3 days will be crossed by a proton and within 30 days by an alpha particle (Curtis S.V. et al., 2000)



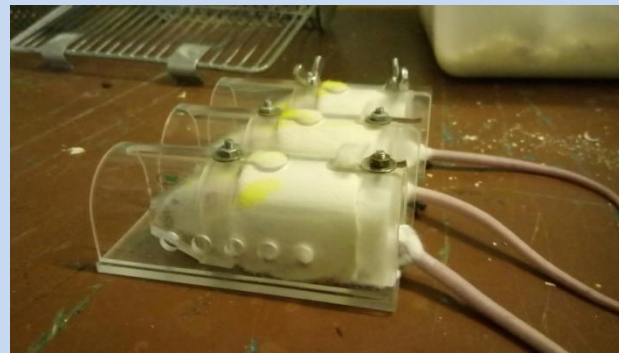
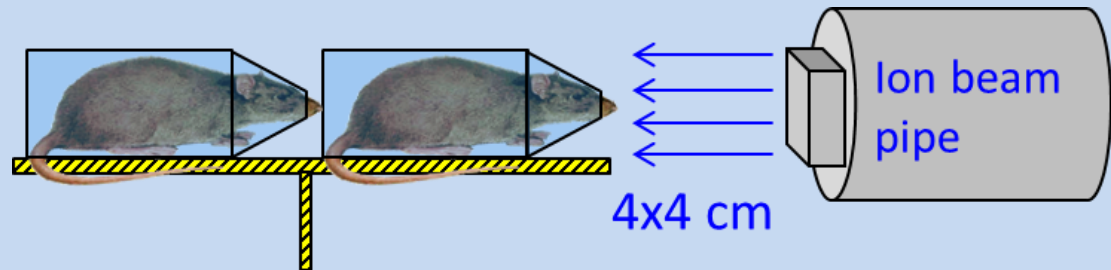
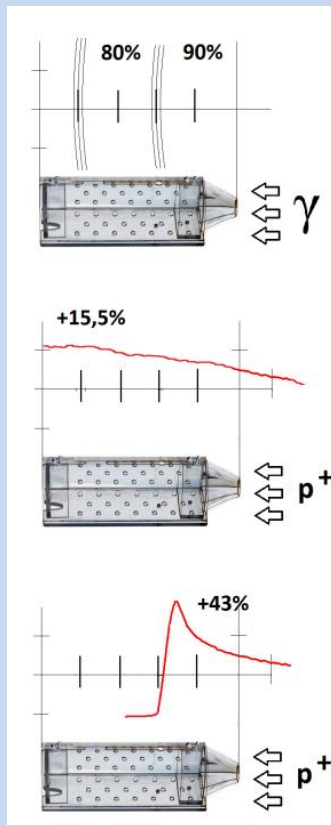
- damage to the glutamatergic transmission in the hippocampal synaptosomes; a significant decrease in the expression of the NR1, NR2A and NR2B subunits of the glutamatergic NMDA receptor





Radiobiological Experiments

- heavy ion-induced functional and morphological disorders in the CNS
- neurochemical studies of the brain neurotransmitter metabolism in rats after exposure to ^{12}C ions and γ -rays
- molecular and genomic damage under irradiation
- research on retina damage in mice by accelerated charged particles





Gamma irradiation

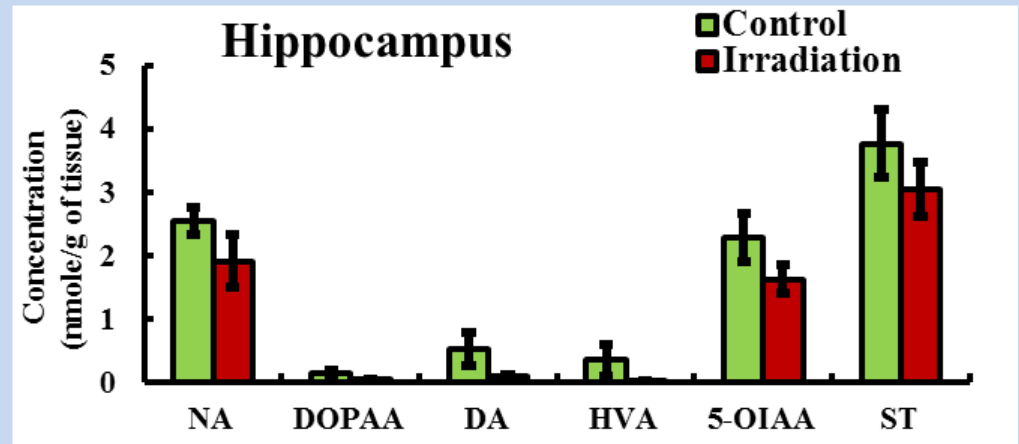
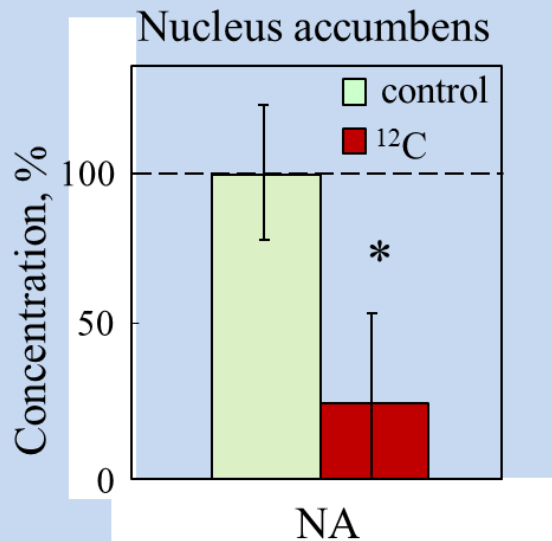
Proton irradiation



Irradiation with 1 Gy of 500 MeV/u carbon ions

- Radiation-induced decrease in the level of neurotransmitters is observed in the brain regions responsible for the emotional and motivational state (radiobiological experiments at the Nuclotron-M; effect of accelerated heavy ions on the functional activity of the CNS)

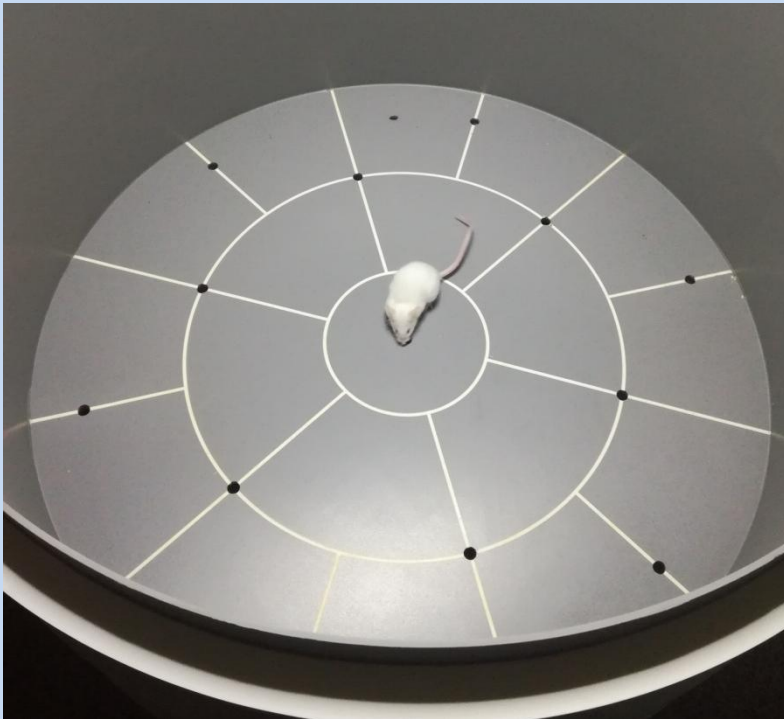
3 months after irradiation





Cognitive-behavioral tests

Open Field

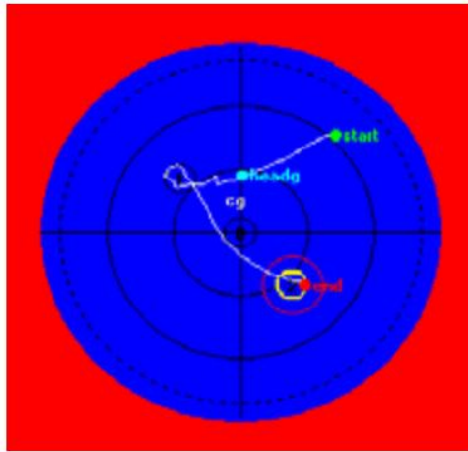


Barnes Maze

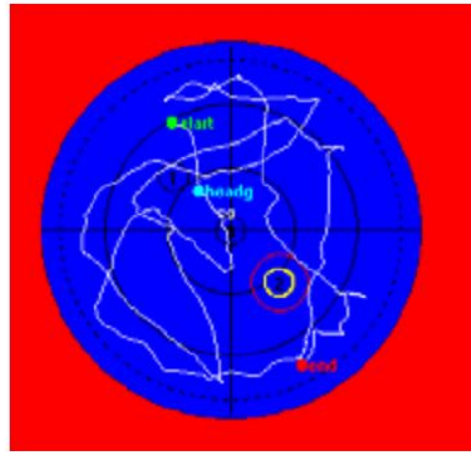


Morris water maze test

Control



1.5 Gy ^{56}Fe Particles



^{56}Fe ions, 1 GeV/amu

1,5 Gy

1 month after irradiation

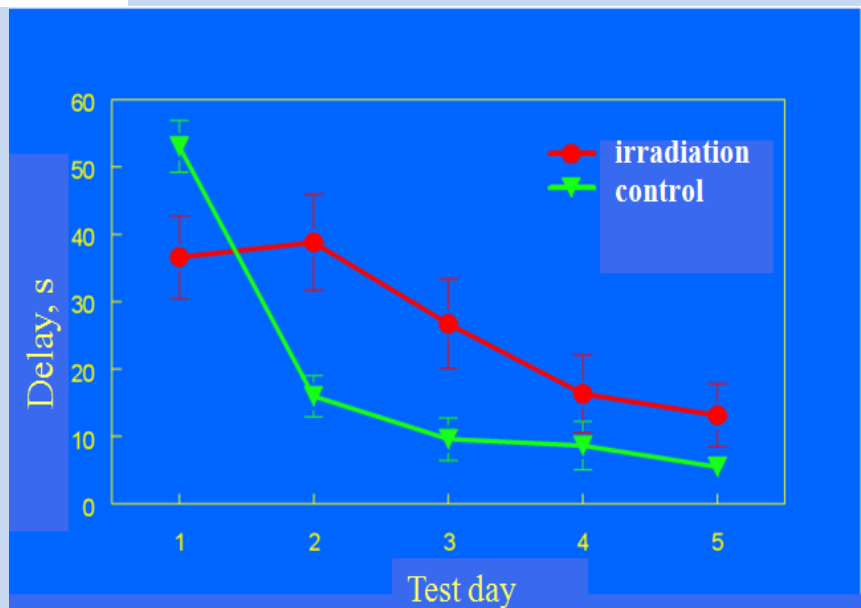
M.Rabin, 2005

Rat 214-126
Morris Water Maze
Learning Test #1

Tracking with:
Noldus Ethovision

(c) Jean-Etienne Poirrier, 2006
Cyclotron Research Center
University of Liege

jepoirrier@ulg.ac.be
<http://www.poirrier.be/~jean-etienne/>



The effect of 1 Gy ^{12}C particle radiation exposure on rats

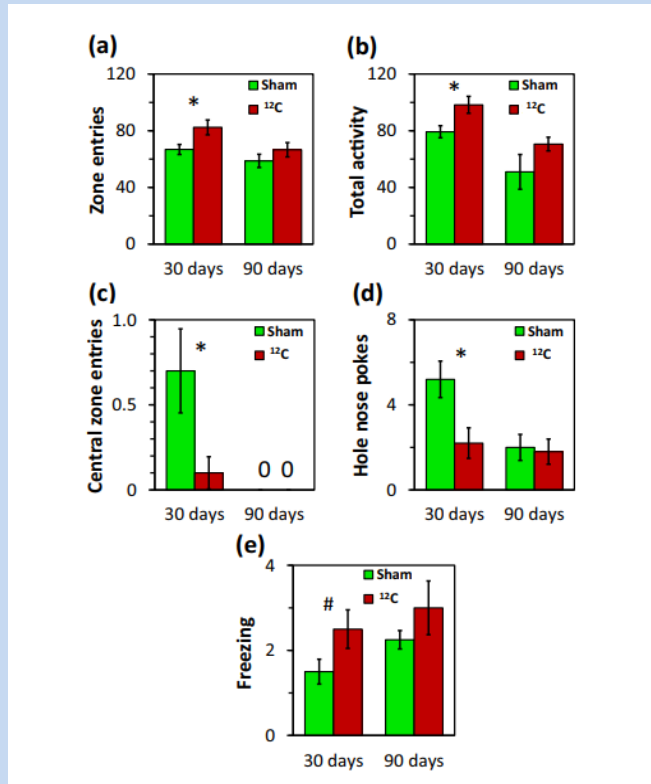


Fig. 1 The open field test results - including zone entries (a), total activity (b), central zone entries (c), hole nose pokes (d), and freezing frequency (e). (*) $p < 0.05$ and (#) $p < 0.1$ between the exposed and sham-irradiated rats at the same periods of time after exposure

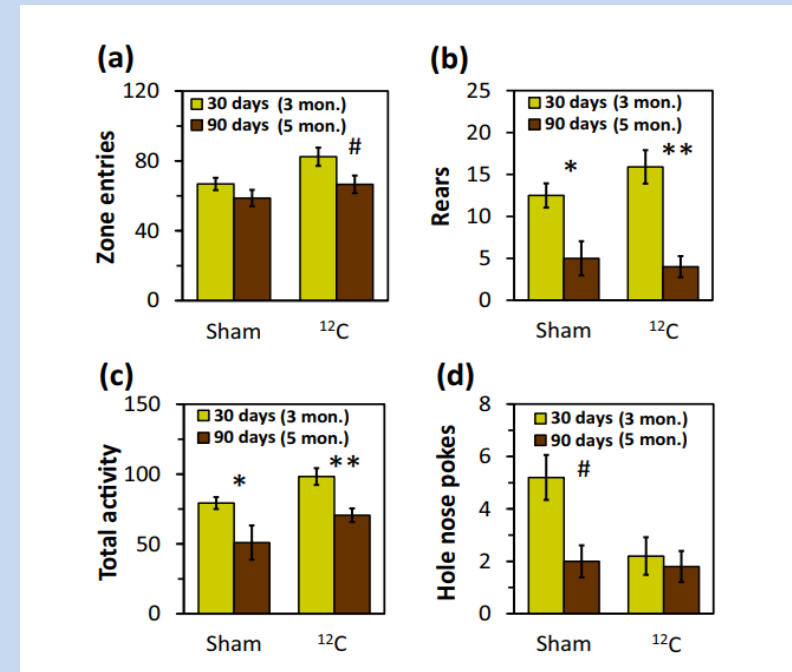


Fig. 2. The temporal dynamics of the open field test - including zone entries (a), rearings (b), total activity (c), and hole nose pokes (d). (*) $p < 0.05$, (**) $p < 0.01$, and (#) $p < 0.1$ between 3- and 5-month-old rats in the exposed and sham-irradiated groups. The corresponding periods after exposure were 30 and 90 days, respectively.

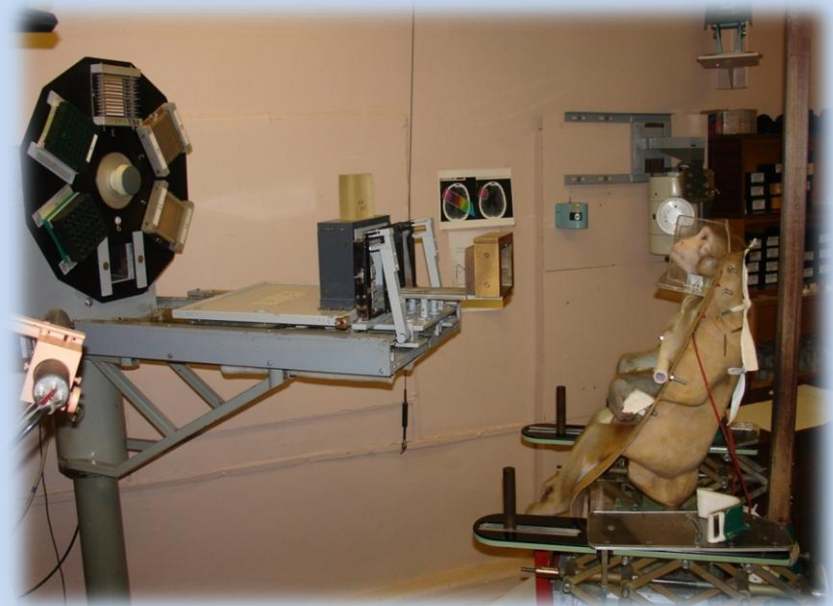


Experiments with monkeys



Proton irradiation with
medical beam, 170 MeV, 3
Gy

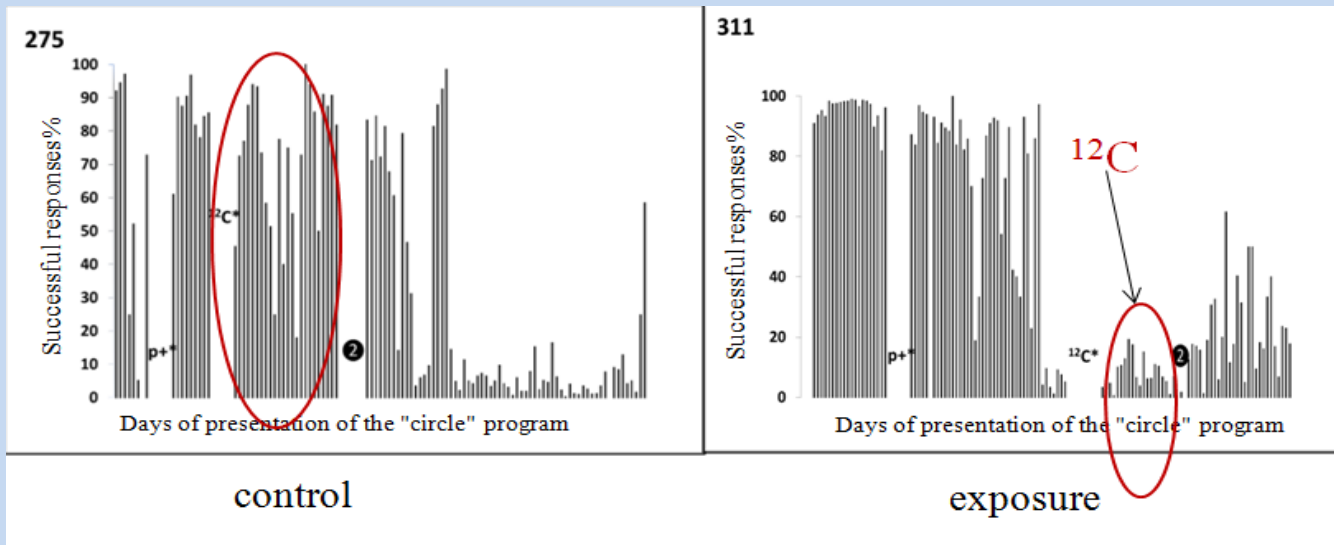
Irradiation with ^{12}C ions, 500 MeV/u, at the
Nuclotron, 1 Gy



Macaca mulatta irradiation (^{12}C ions, 500 MeV/nucleon)

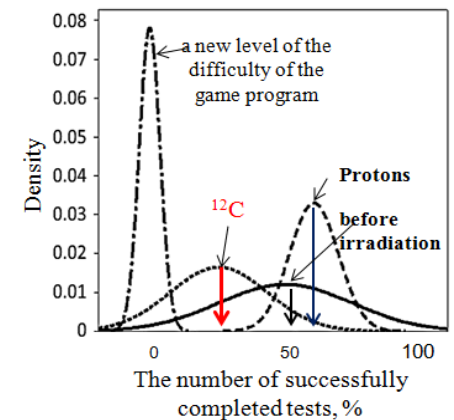
Psychological Test System — a series of 18 computer gaming tasks of increasing difficulty to simulate the basic elements of the operator's activity

Indicators of cognitive functions in the irradiated and control monkey groups

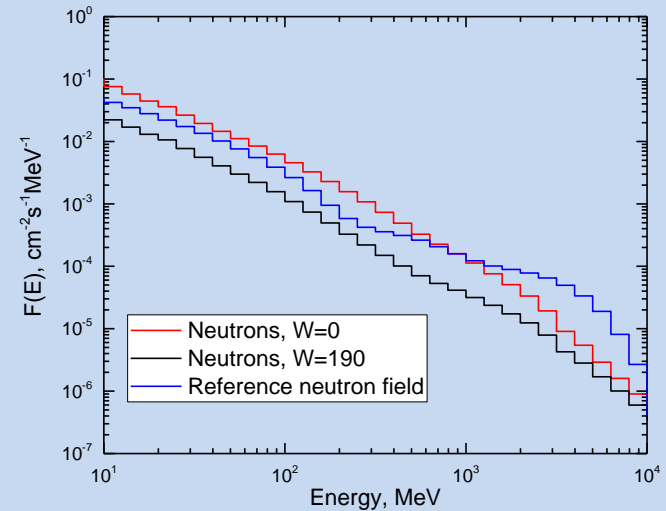
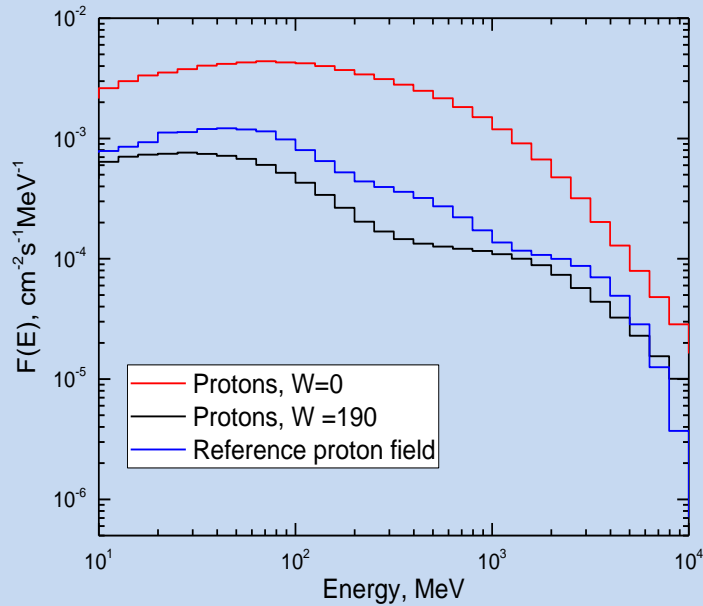


Dose: 1 Gy

p + — proton irradiation day; ^{12}C — carbon ion irradiation day; 2 — a new level of the game program difficulty.



NUCLOTRON BASED SIMULATION OF RADIATION ENVIRONMENT ON THE BOARD OF SPACECRAFT IN DEEP SPACE



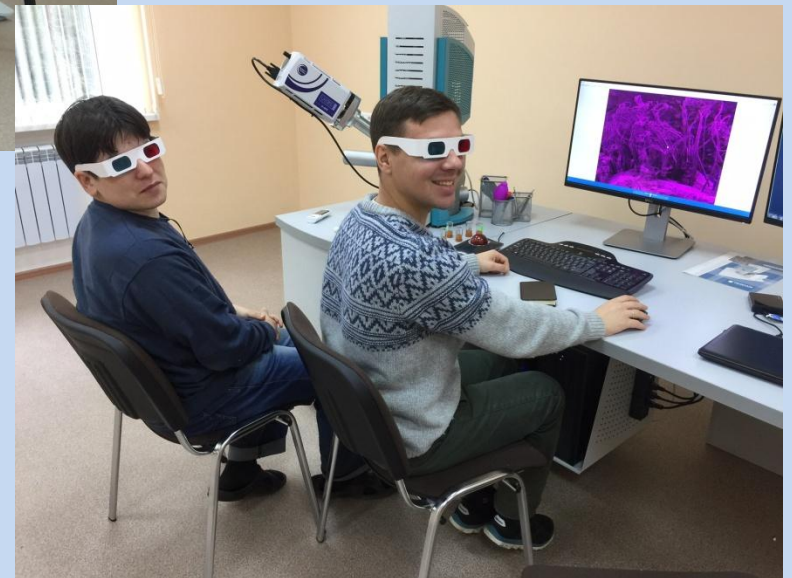
Modeling the actual conditions of radiation in space with accelerated-based radiation experiments

The methods based on the linear combination of energy spectra of the particles emitting at various angles from three different targets, consecutively bombarded by 10 GeV proton beam.



Group of Astrobiology

SEM Tescan Microscope



Thank you for your attention!

