



DUBNA



JINR

Laboratory of Radiation Biology

LRB structure

Department of Radiation Biology and Physiology:

- ❑ Molecular Radiobiology Sector;
- ❑ Radiation Cytology Sector;
- ❑ Radiation Physiology Sector;
- ❑ Radiation Neurochemistry Sector;
- ❑ Mathematical Modeling Sector;
- ❑ Lower Eukaryote Radiation Genetics Group.

Department of Radiation Research:

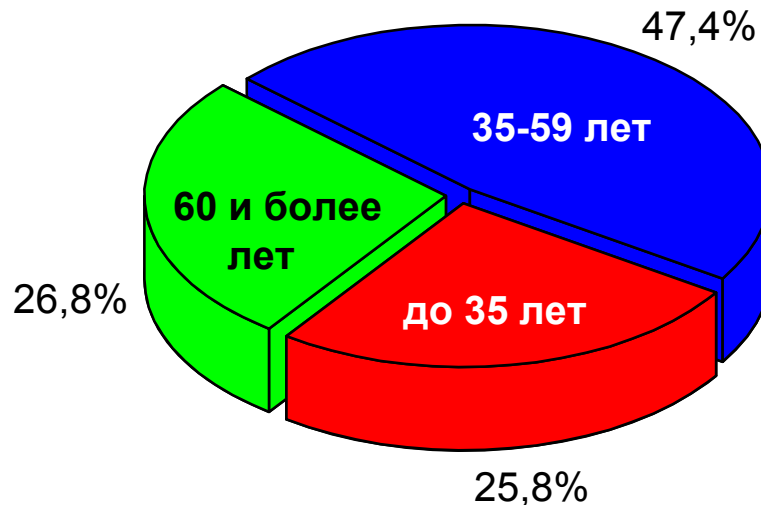
- ❑ Group for Modeling Ionizing Radiation Interaction with Matter;
- ❑ Group for Studying Radiation Fields of JINR's Basic Facilities and Environment.

❖ *Astrobiology Sector*

The LRB's full-time staff is 101

The LRB staff includes 3 Members of the Russian Academy of Sciences (RAS), 6 Doctors of Biological Sciences, 2 Doctors of Medical Sciences, 2 Doctors of Physical and Mathematical Sciences, 9 Candidates of Biological Sciences, and 8 Candidates of Physical and Mathematical Sciences.

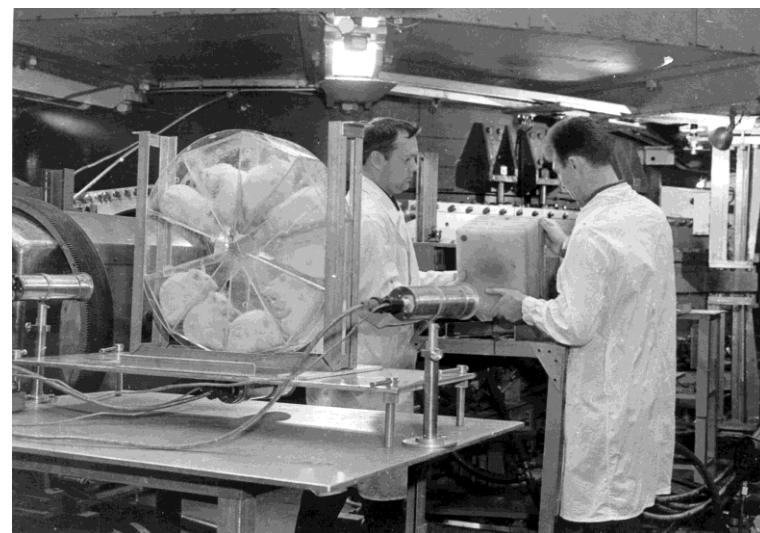
Age distribution:



First radiobiological experiments at the Phasotron



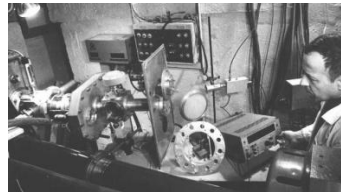
Protons with
energy up to 660
MeV



Laboratory of Radiation Biology

- ❑ **1978** Sector of biological research in LNP
- ❑ **1988** Division of biophysics in LNP
- ❑ **1995** Department of Radiation and Radiobiological Research of JINR

- ❑ **2005** *Laboratory of Radiation Biology*



The main field of research:

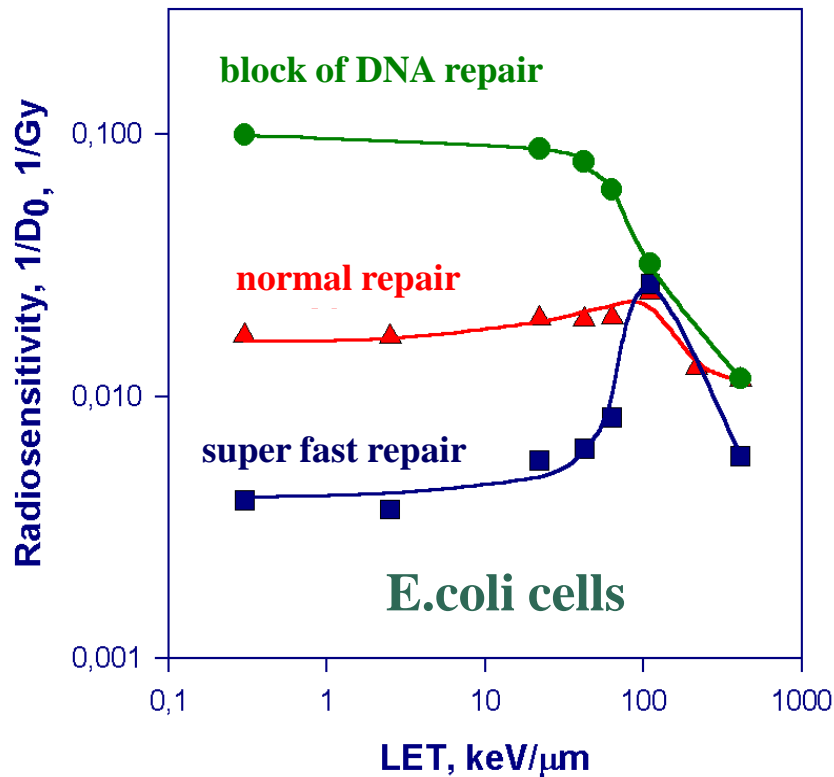
THE BIOLOGICAL ACTION
OF HEAVY CHARGED
PARTICLES OF DIFFERENT
ENERGIES



What radiobiological problems
can be solved at use of accelerated
heavy particles?

A.
Heavy ions is a powerful tool
for solving problems in
radiation genetics

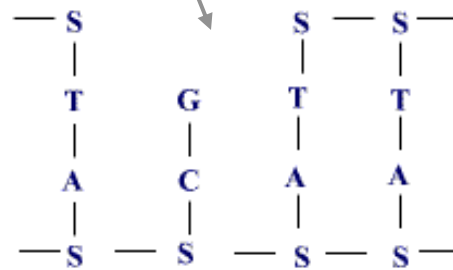
The RBE problem



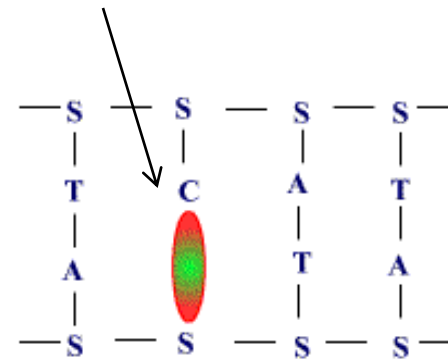
DNA repair capacity of the living cells determines the type of RBE on LET dependence

Single DNA damage

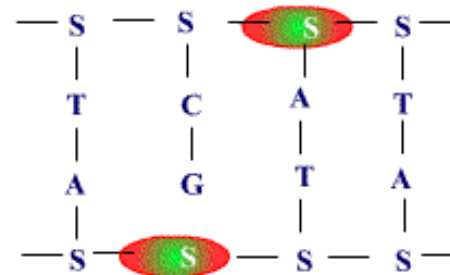
Single strand break



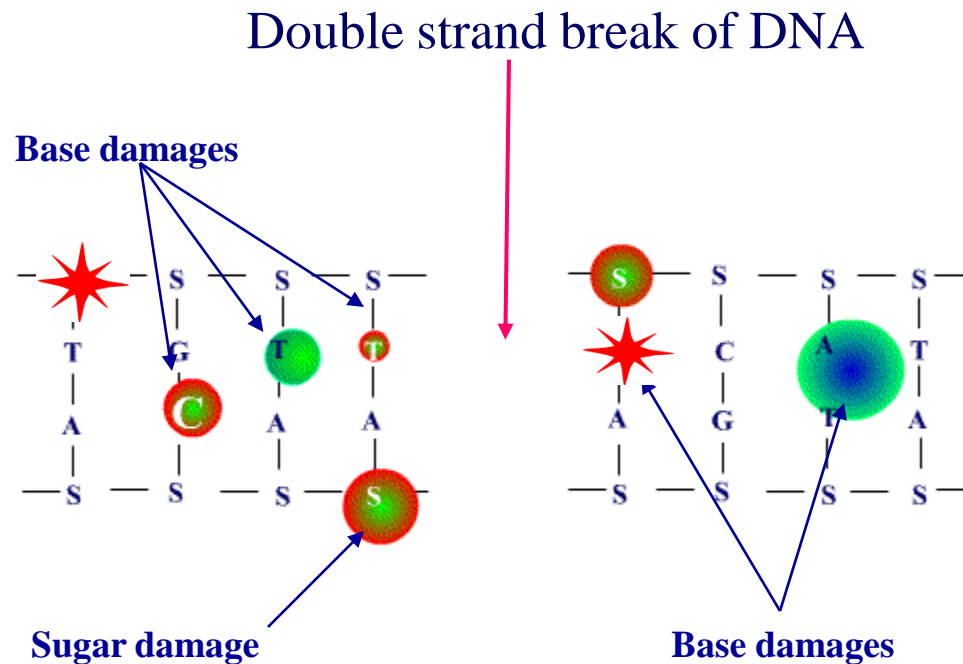
Base damage



Sugar damage

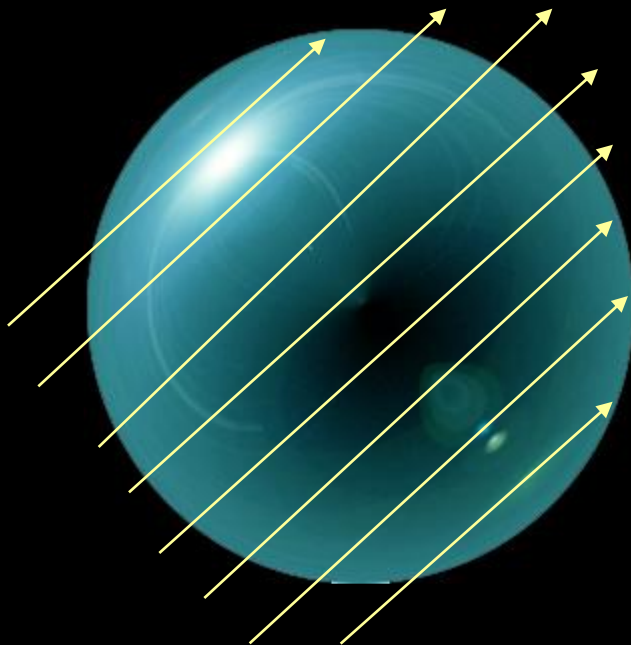


Clustered DNA damage



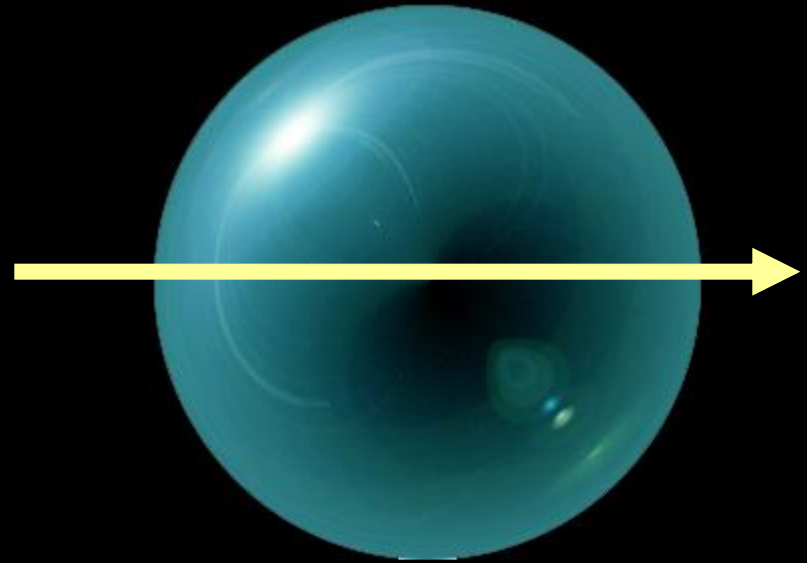
Radiation dose distribution in matter

1 unit of the dose



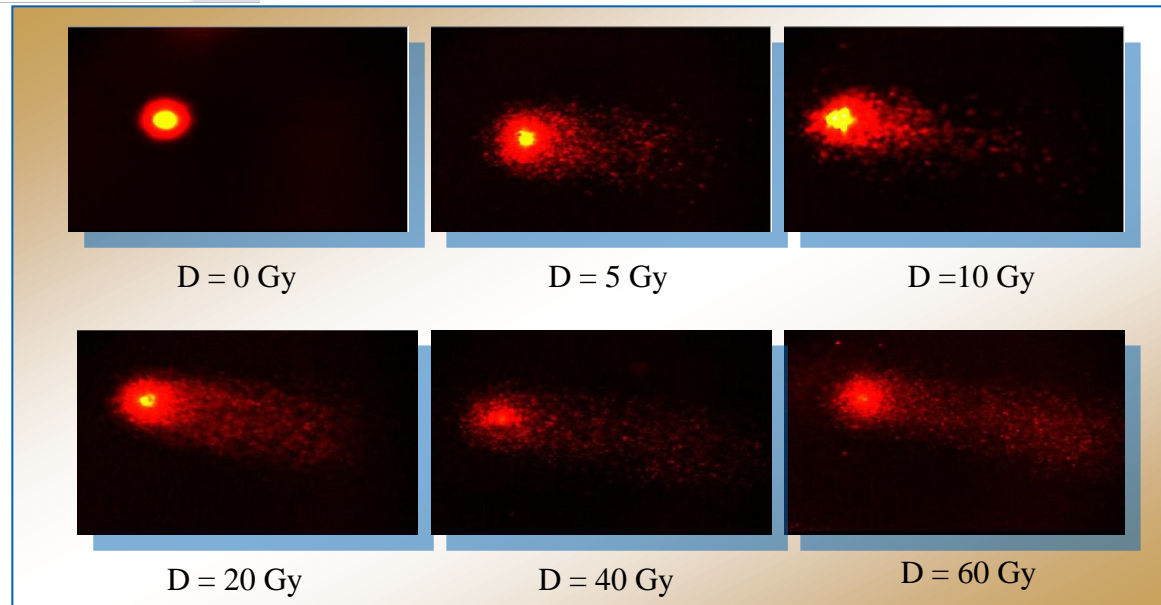
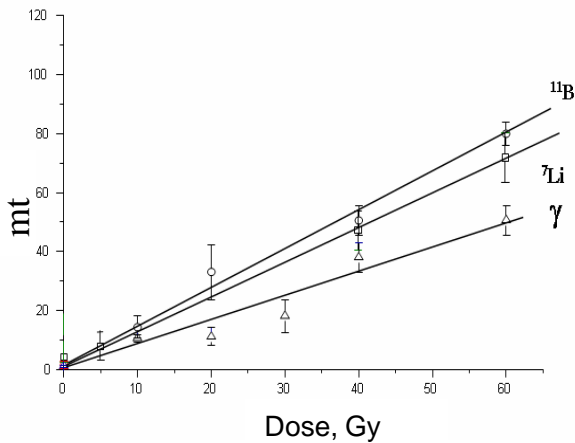
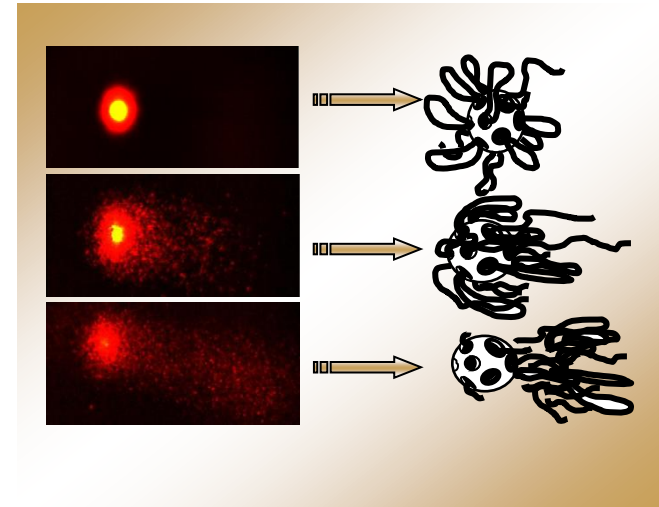
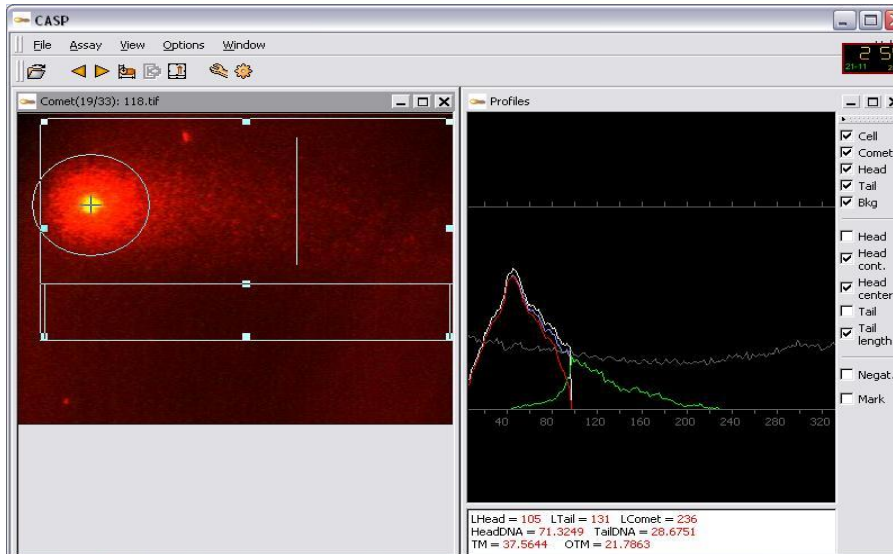
X-rays

1 unit of the dose



Fe ion

“Comet assay” for detection of DNA lesions

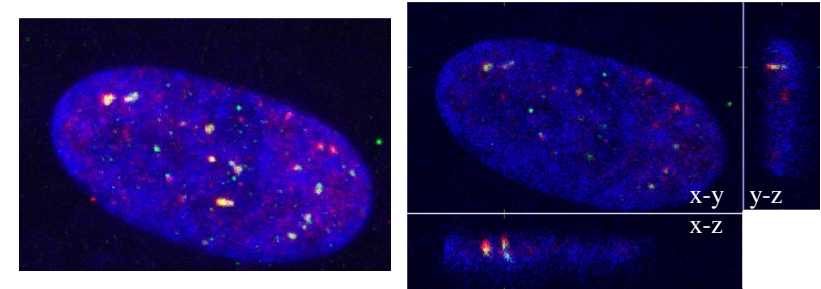
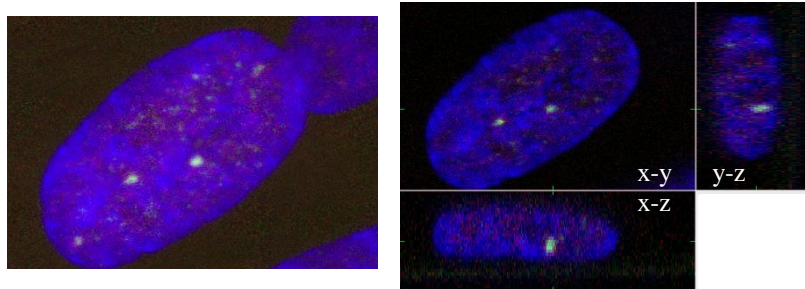


Human cells exposed to γ -rays and ^{11}B ions

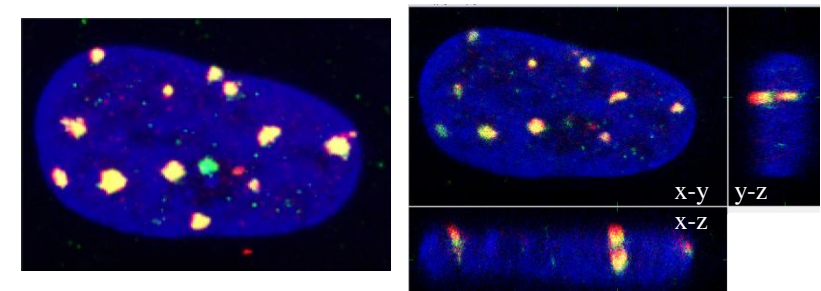
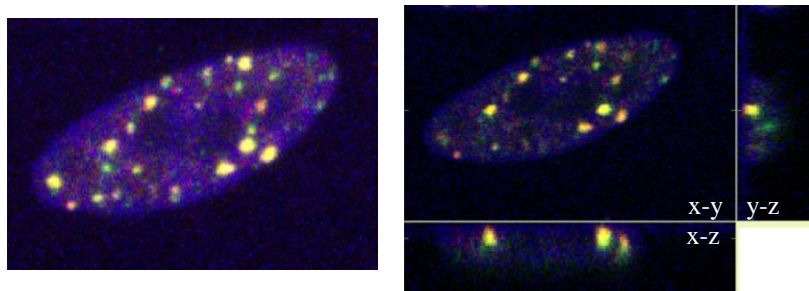
γ -rays (LET = 0.3 keV/ μm)

^{11}B ions (LET = 135 keV/ μm)

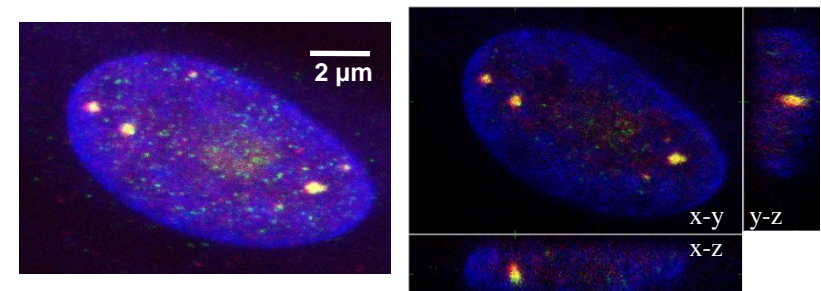
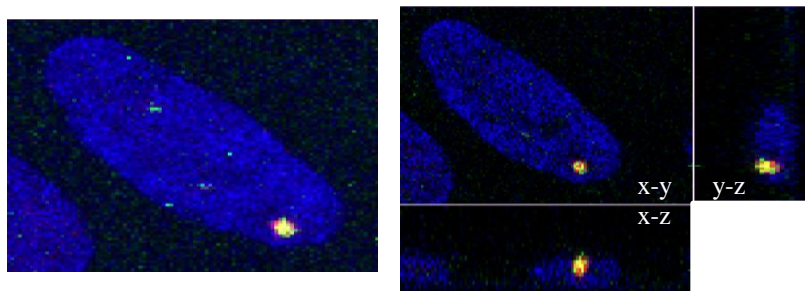
5 min



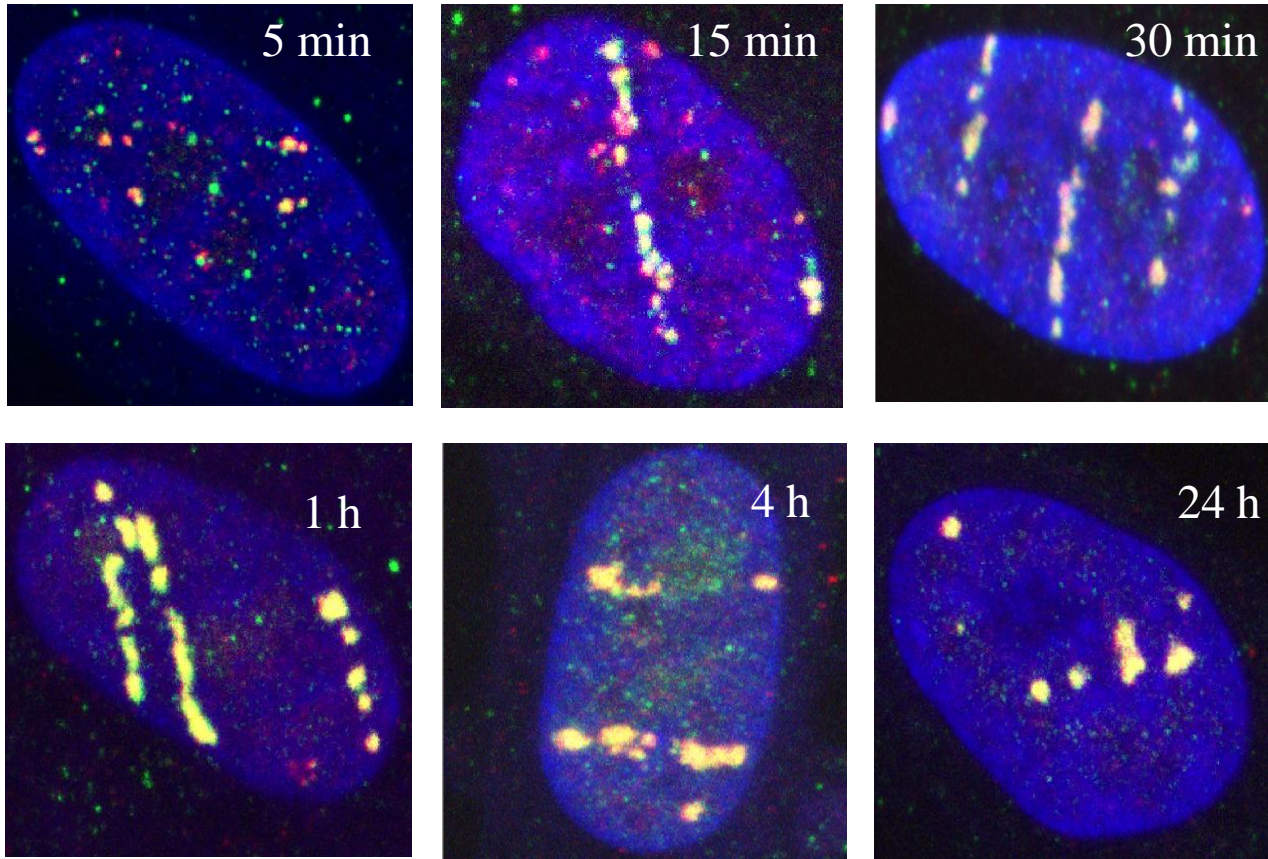
1 h



24 h



Human cells exposed to ^{11}B ions at 10°

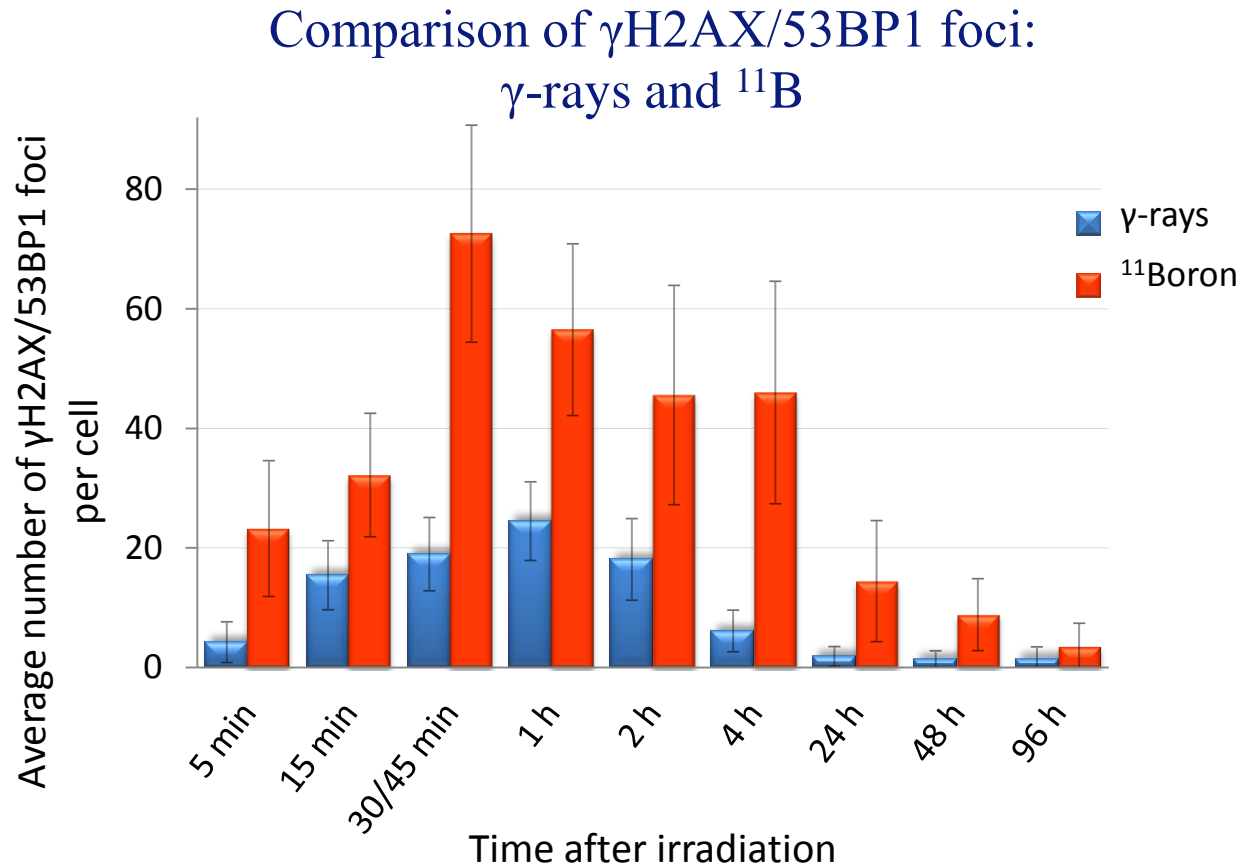


● γH2AX ● 53BP1 ● Chromatin (DAPI)

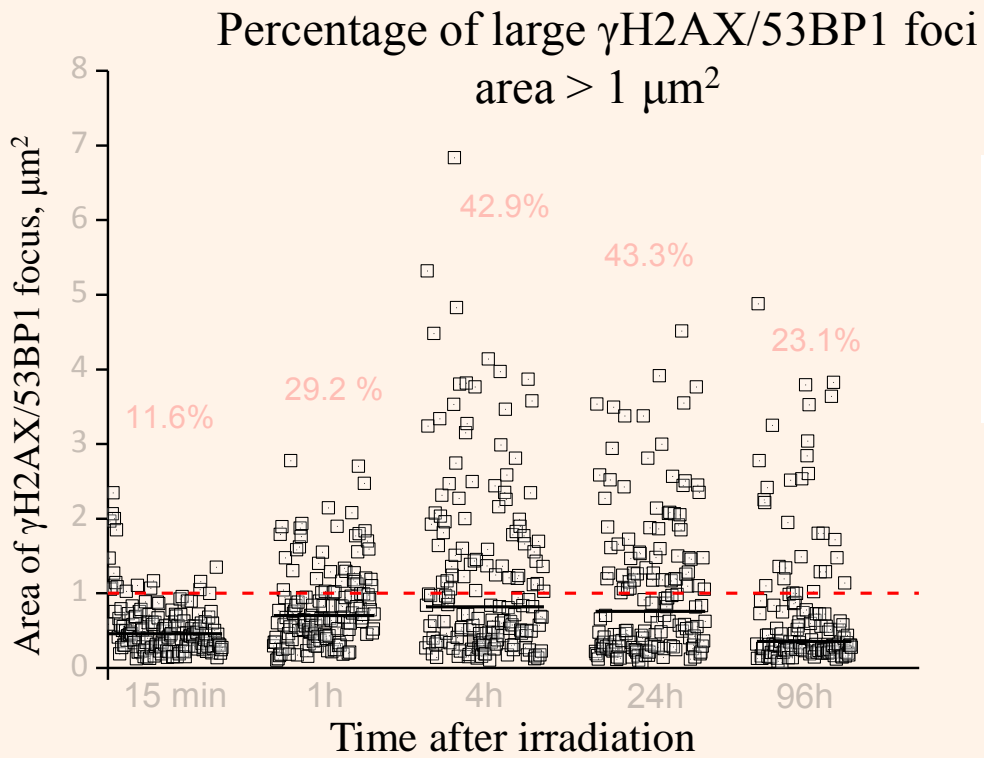
3D visualization of DNA foci streaks



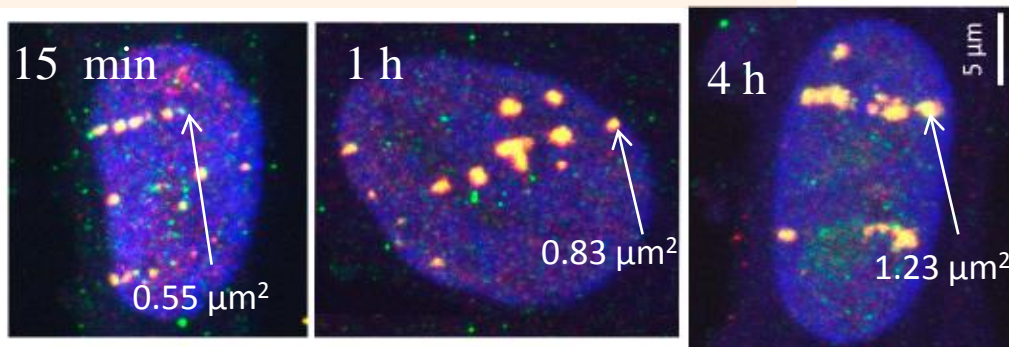
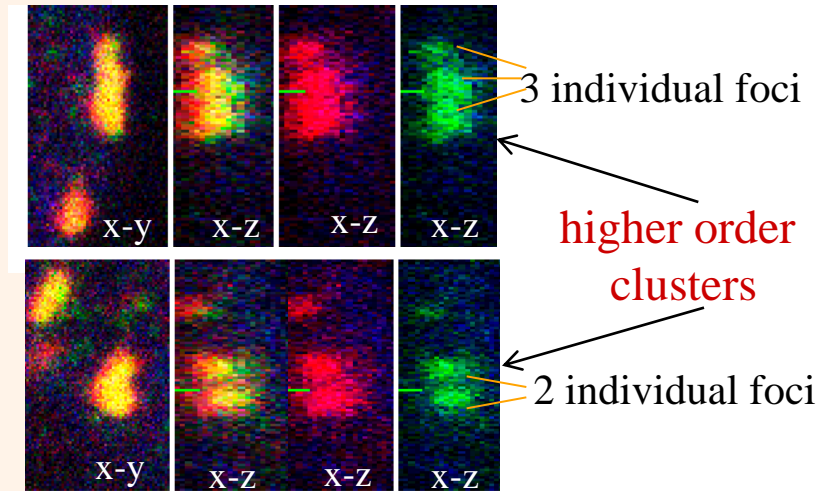
Kinetics of the formation and disappearance of γ H2AX/53BP1 foci



Higher-order clustered γ H2AX/53BP1 foci after ^{11}B irradiation

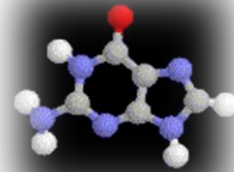


γ H2AX/53BP1 53BP1 γ H2AX

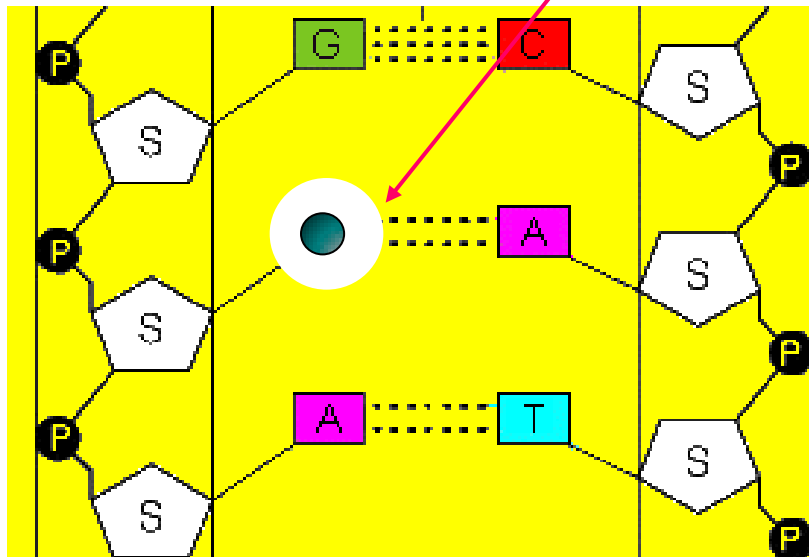


Radiation induced mutagenesis

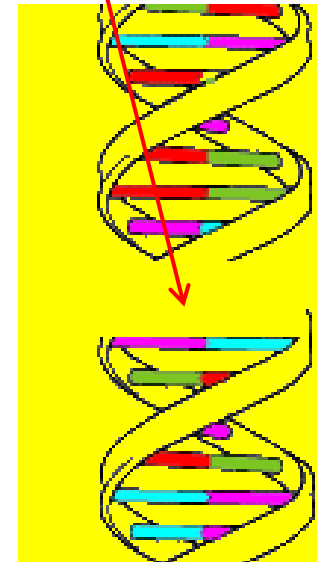
Gene mutation



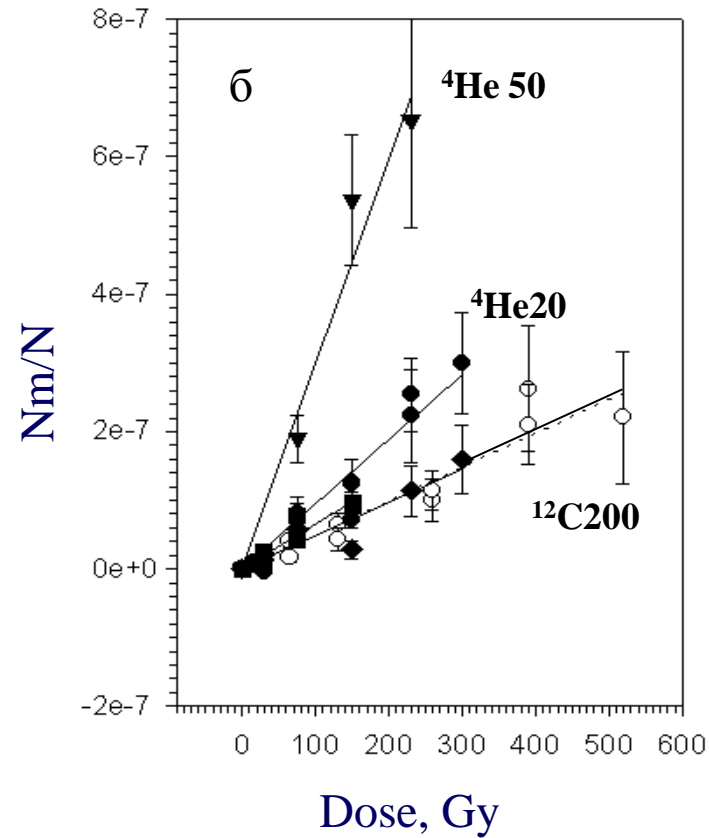
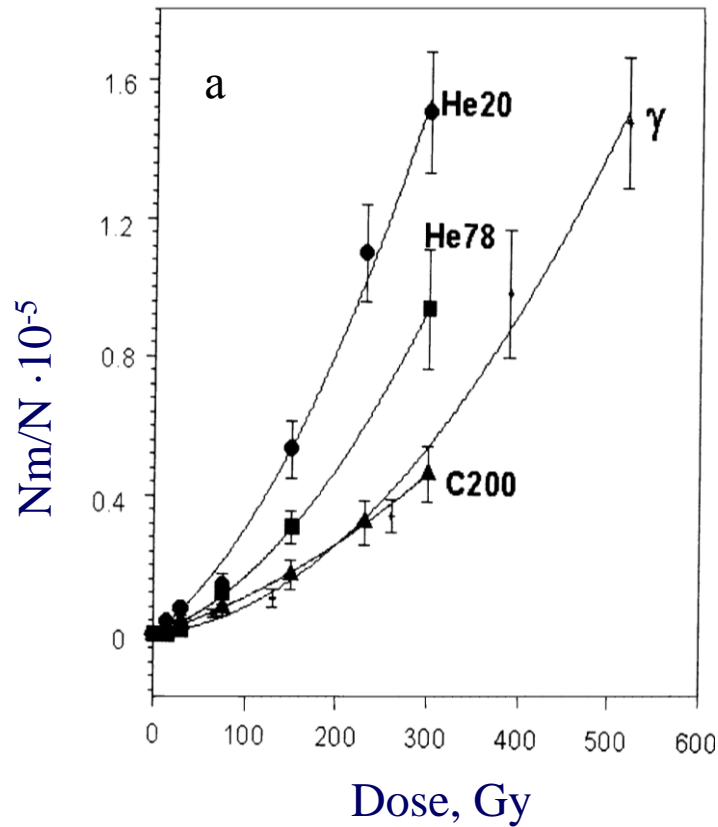
Guanine



Structural mutation

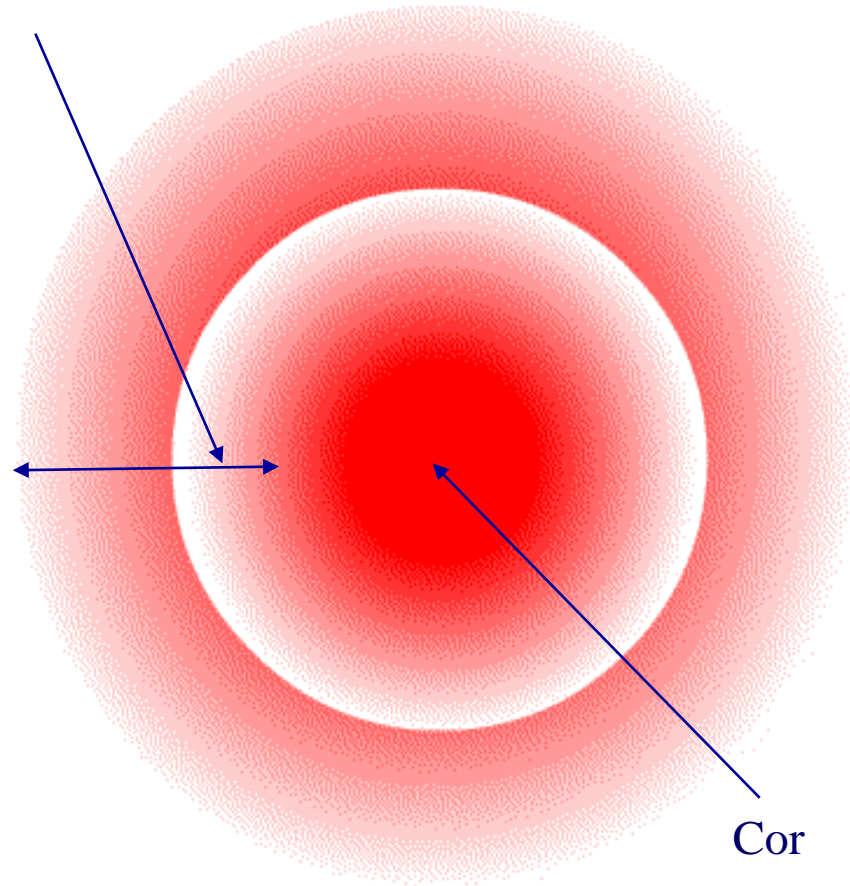


The frequency of gene and structural mutation induction after γ -ray and heavy ion irradiation



Mutagenic belt

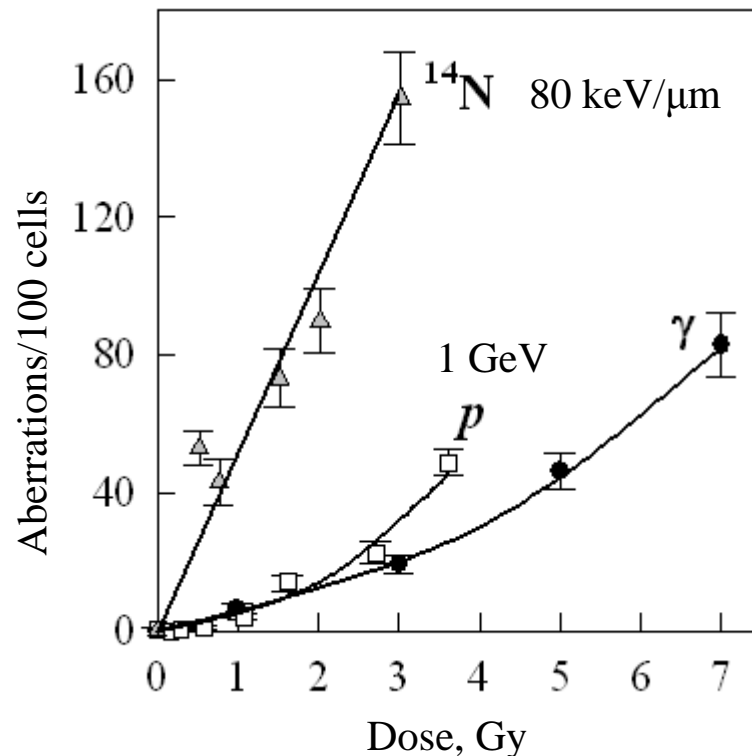
Mutagenic belt of heavy particle track



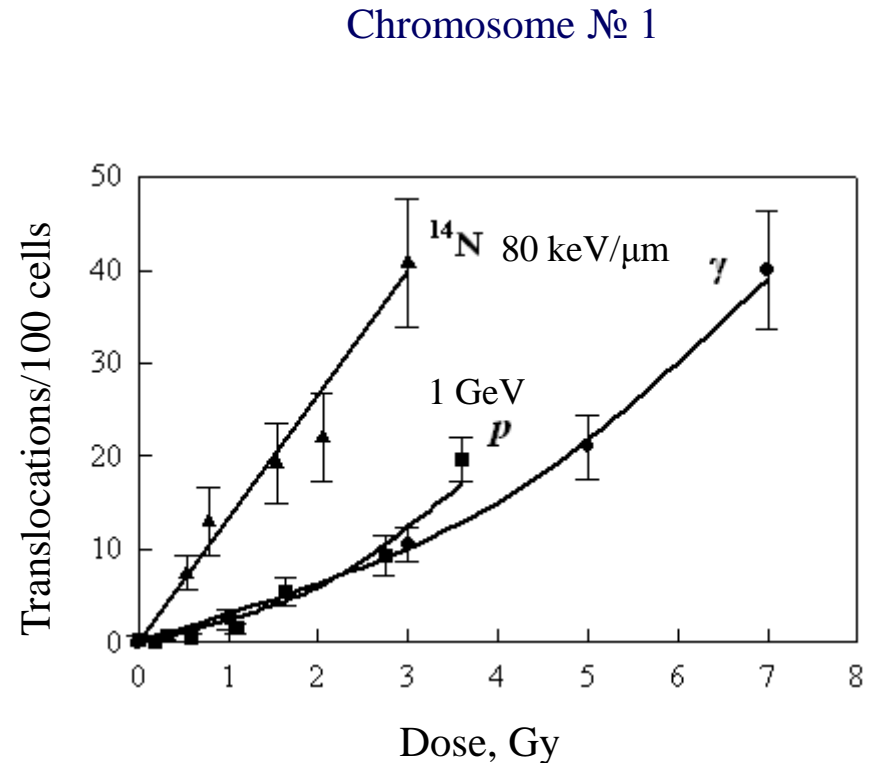
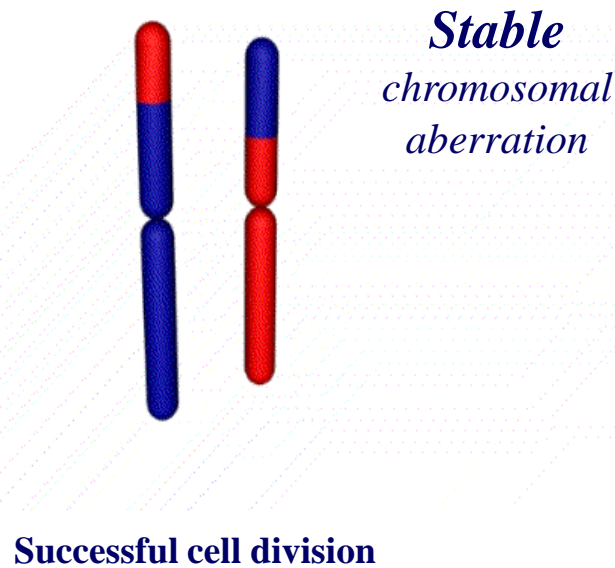
Formation of unstable chromosomal aberration in human cells after heavy ion irradiation

*Unstable
chromosomal
aberration*

Cell division blocking



Formation of stable chromosomal aberrations in human cells after heavy ion irradiation

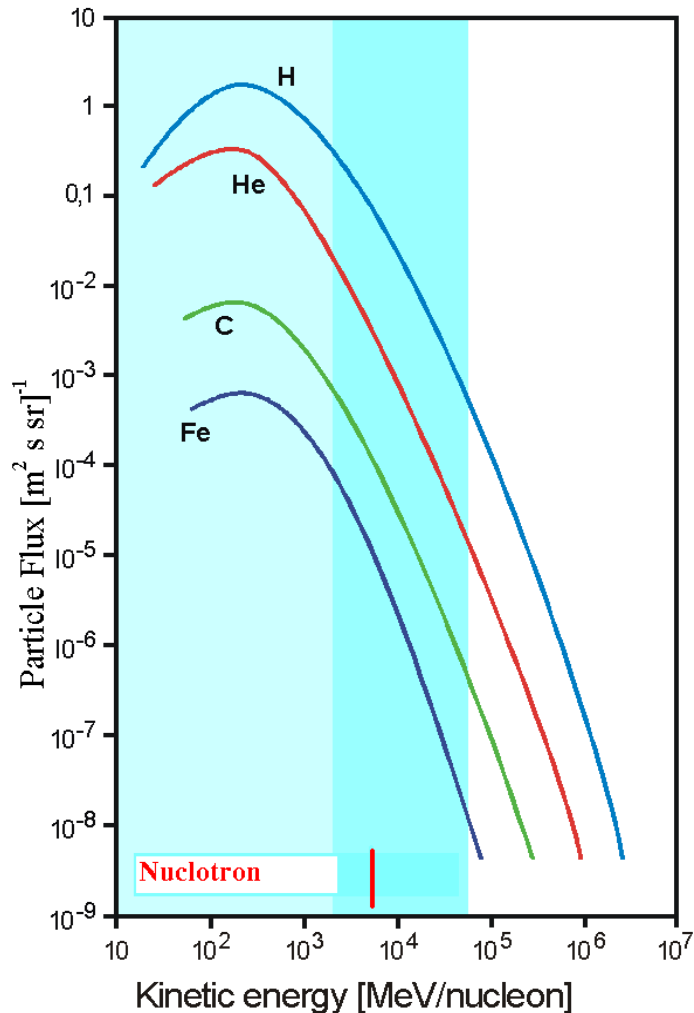




B.

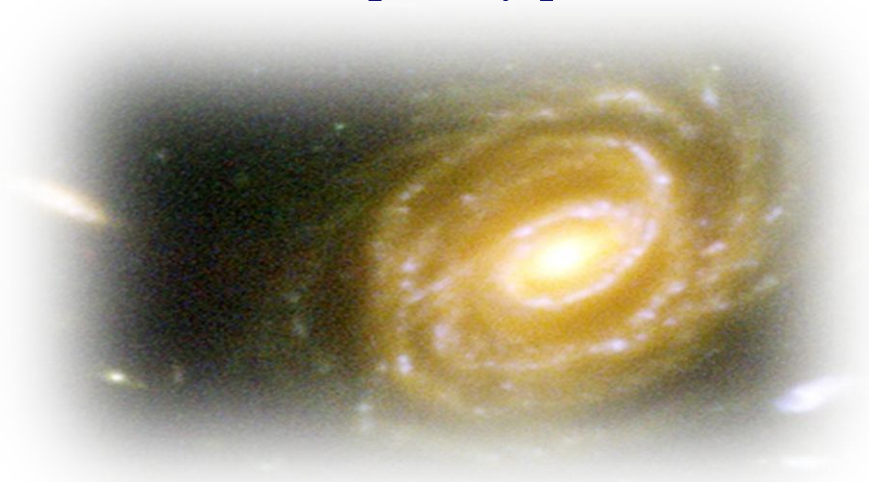
Accelerated heavy ions as a tool
for modeling of biological action
of space radiation

The GCR energy spectrum



The integral flux of GCR particles of carbon and iron groups equals to $\sim 10^5 \text{ part/cm}^2 \text{ per year}$

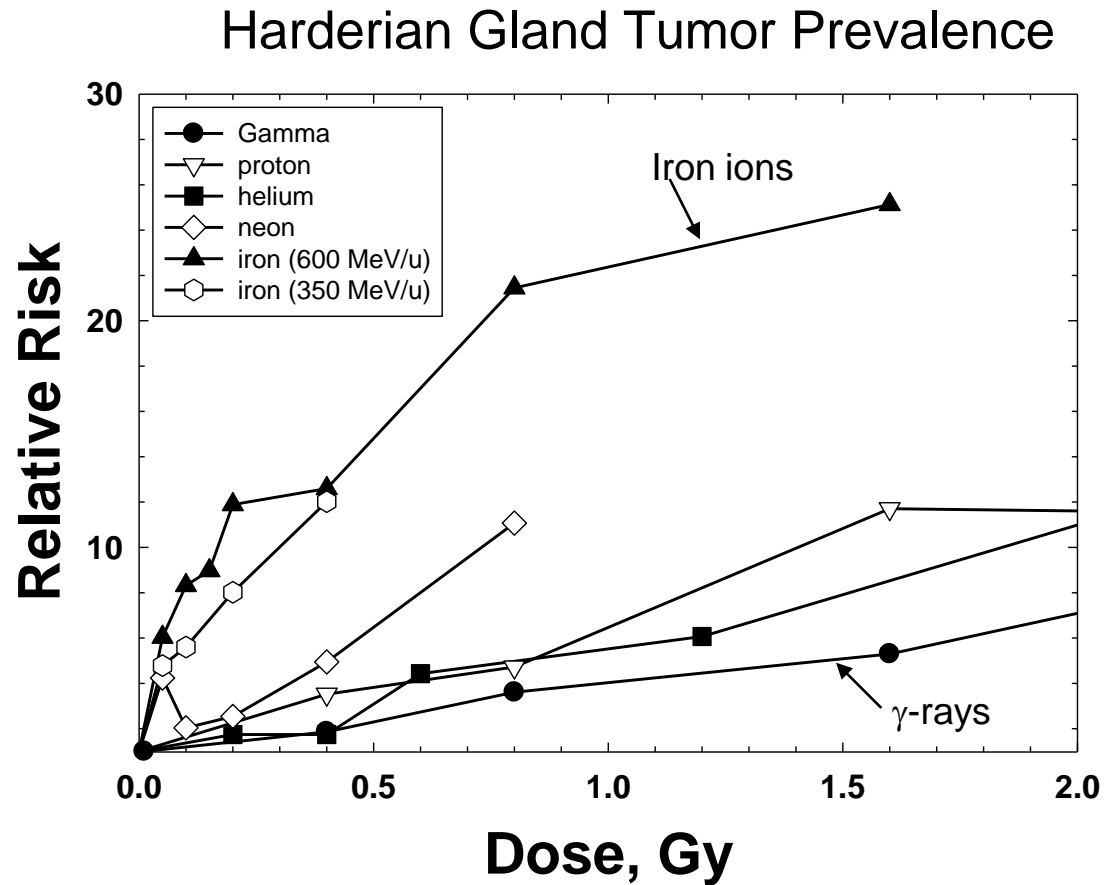
Particle flux density
interplanetary space $Z \geq 20$
160 per day per cm^2

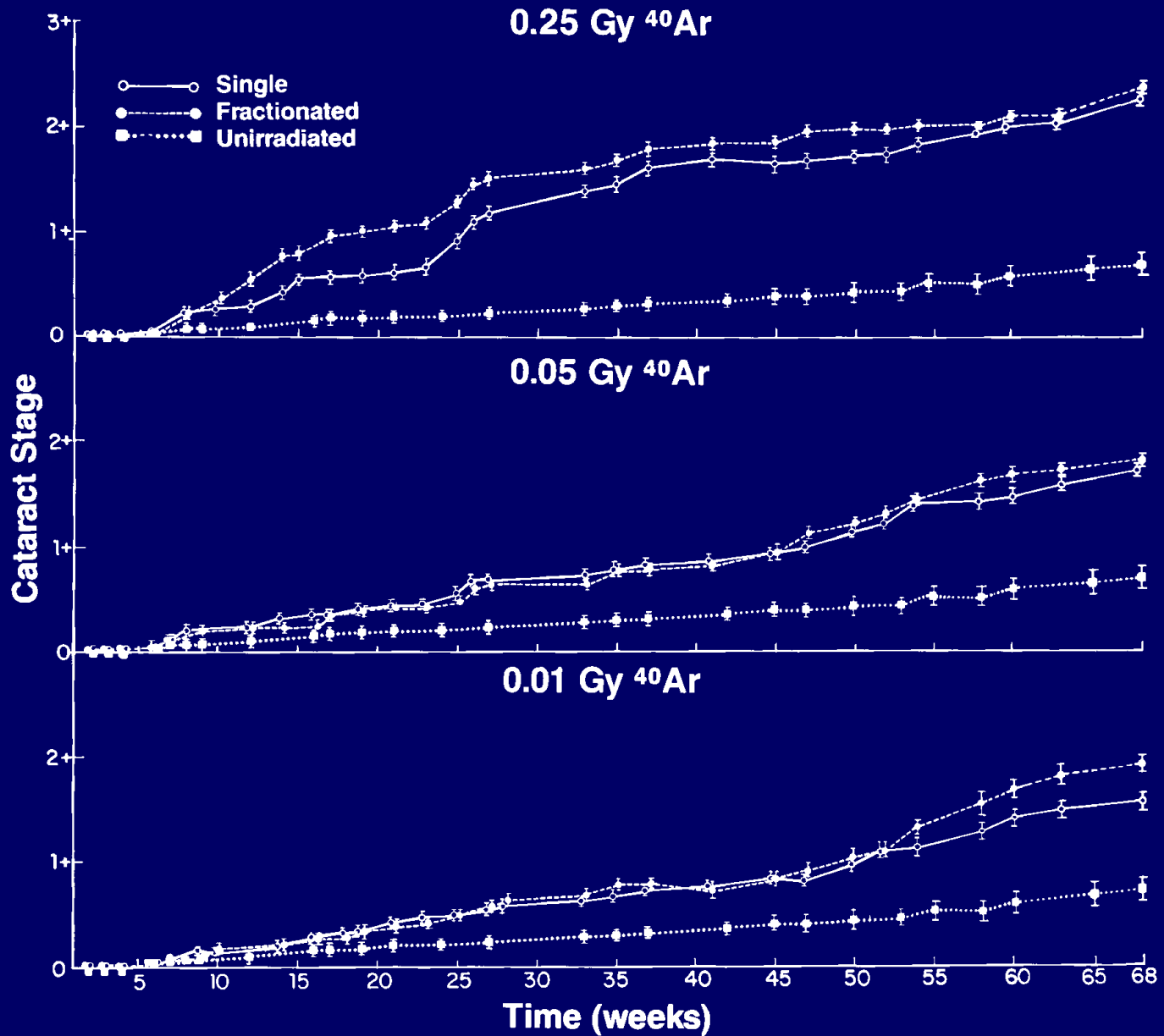


Consequences of Galactic heavy ion action

- ❑ Induction of cancer;
- ❑ Formation of gene and structural mutations;
- ❑ Violation of visual functions:
 - ❑ *lesions of retina;*
 - ❑ *cataract induction;*
 - ❑ CNS violation

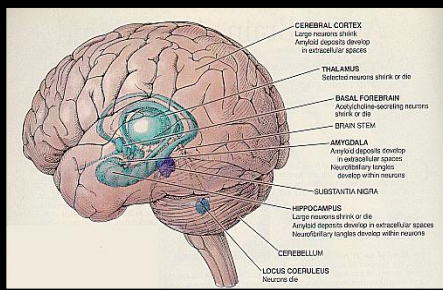
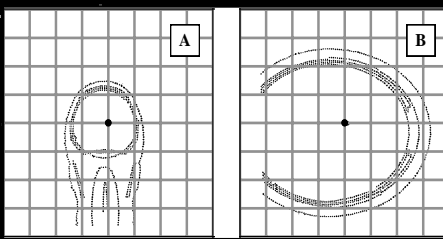
Gardner tumors





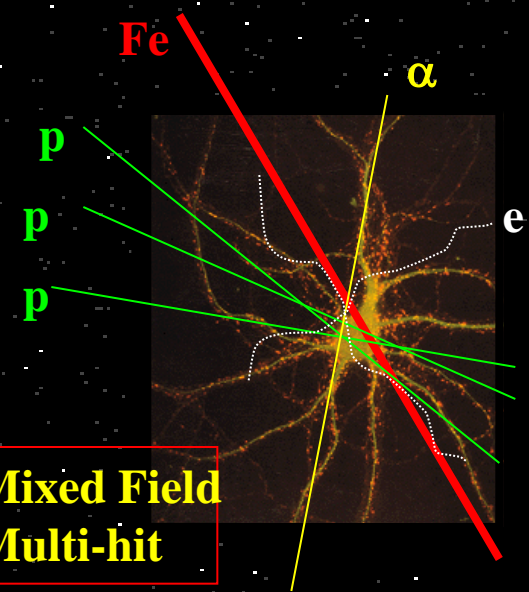
Accelerated heavy ions and CNS

Cosmic ray hit frequencies in CNS critical areas

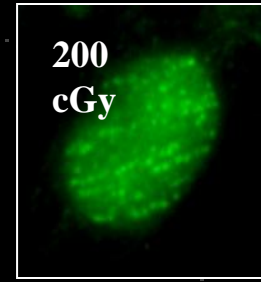
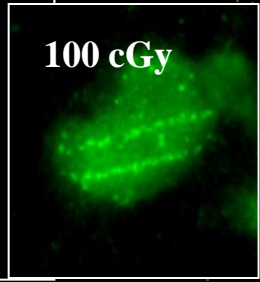
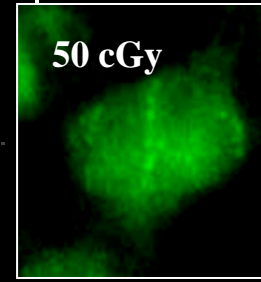
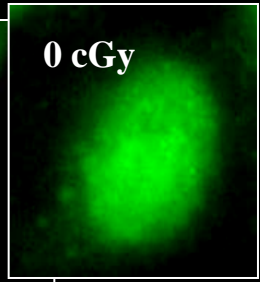
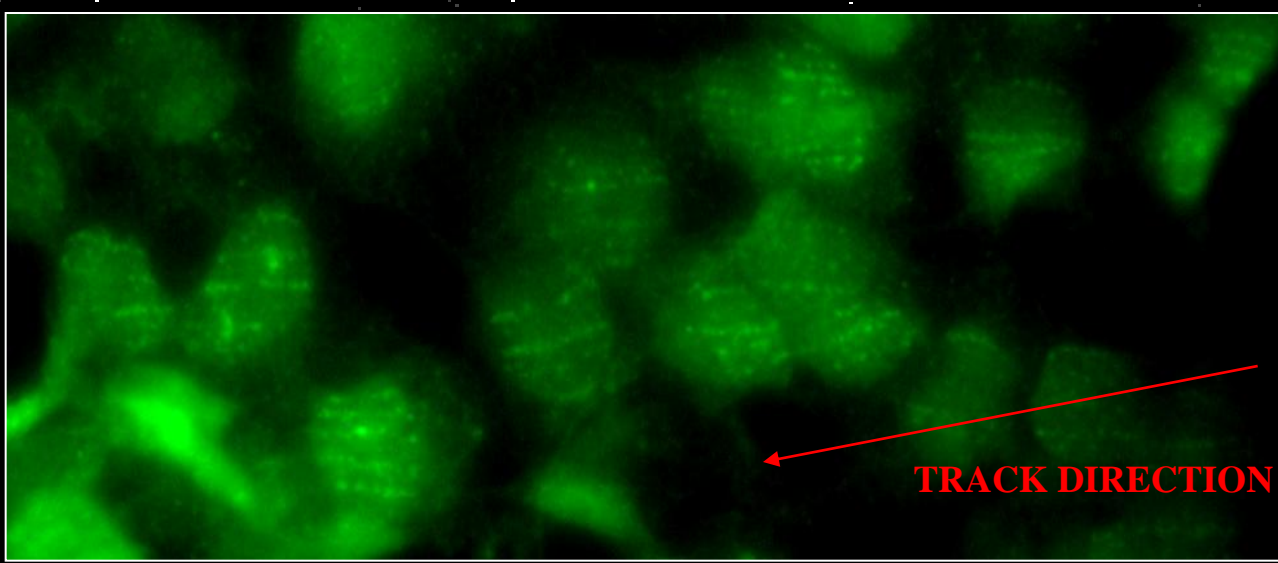


CNS in General

- 2 or 13% cells will be hit at least one Fe particle
- 8 or 46% would be hit by at least one particle with $Z \geq 15$
- Every nucleus will be traversed by a proton once every 3 days and a alpha particle once every 30 days.



FE ION TRACKS VISUALIZED BY MARKERS OF DNA DSBs (γ H2AX)



TRACK DIRECTION

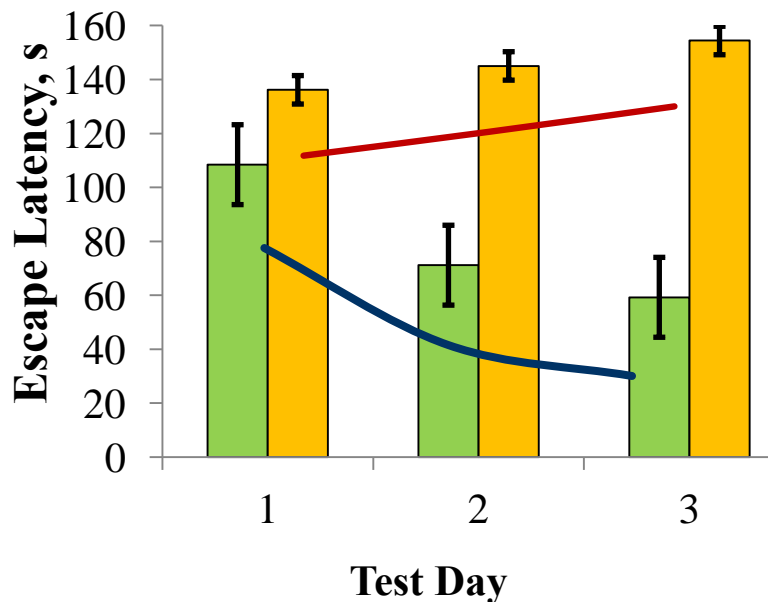
Тест К. Барнс



Behavioral deficits measured 1.5 and 3 months after charged particle exposure

Rats

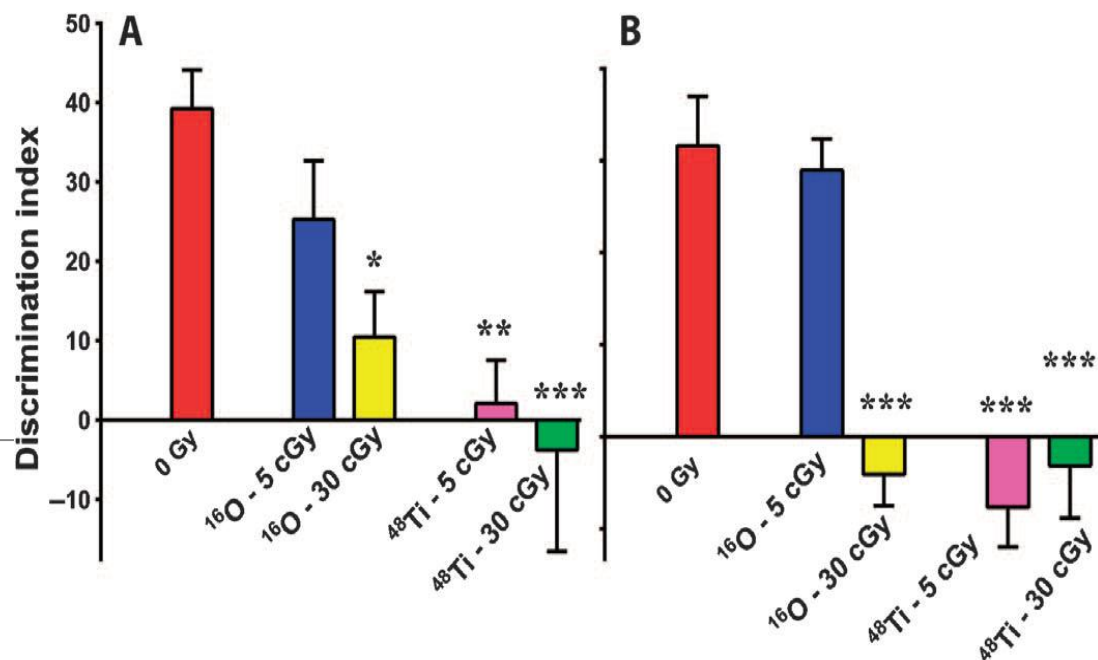
3 months after irradiation



■ control
■ ⁵⁶Fe ions 1 GeV/u, 20 cGy

Mice

6 weeks after irradiation



A - novel object recognition task
 B - object in place task



R. Britten et al., 2012

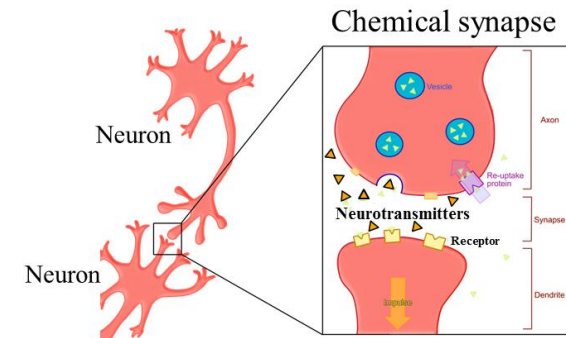
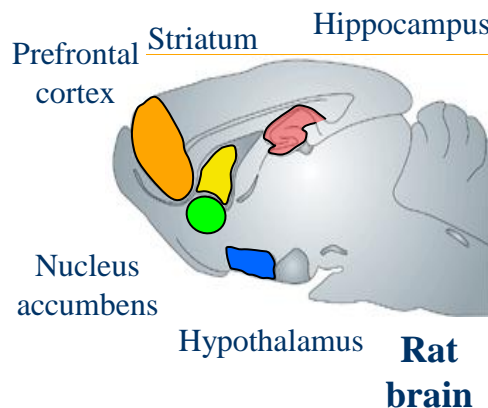
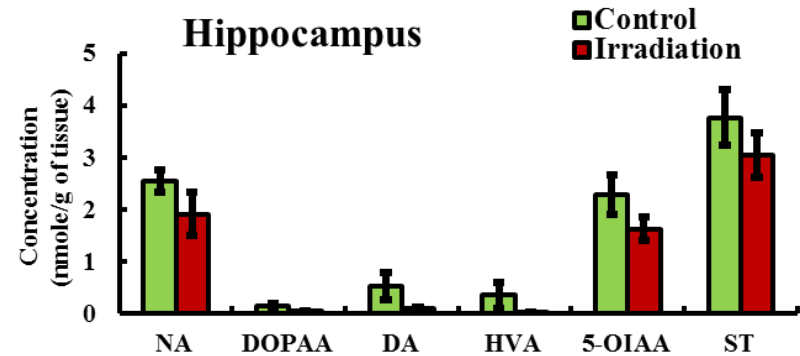
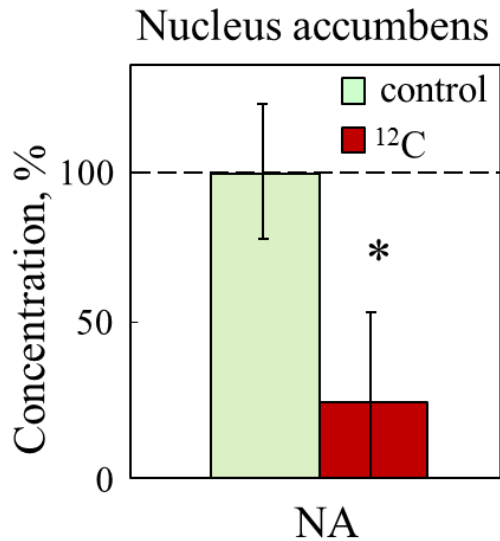
V. Parihar et al., 2015

Studying the level of neurotransmitters in different rat brain areas

Irradiation with 1 Gy of 500 MeV/u carbon ions

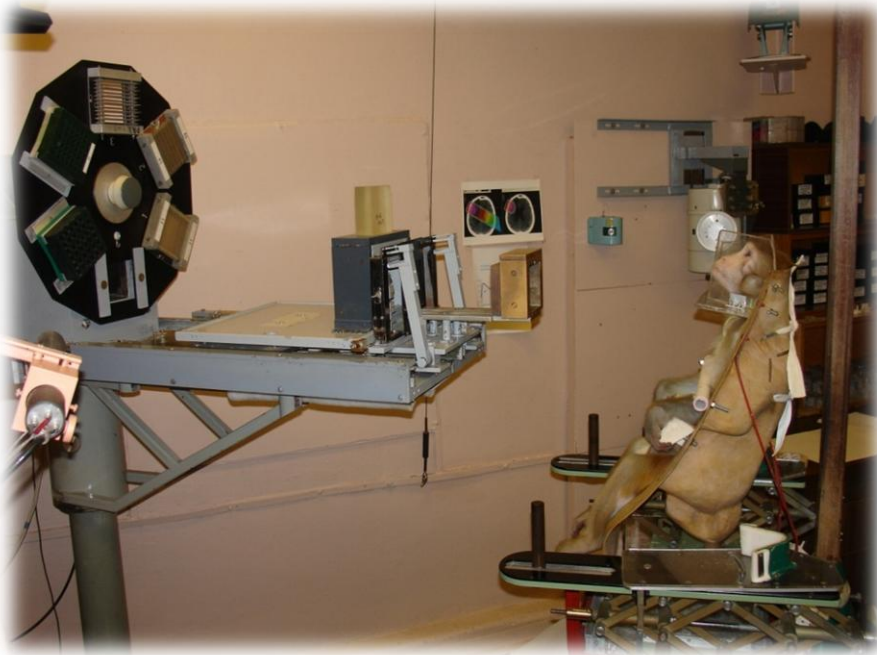
Radiation-induced decrease in the level of neurotransmitters is observed in the brain regions responsible for the *emotional and motivational state*

3 months after irradiation



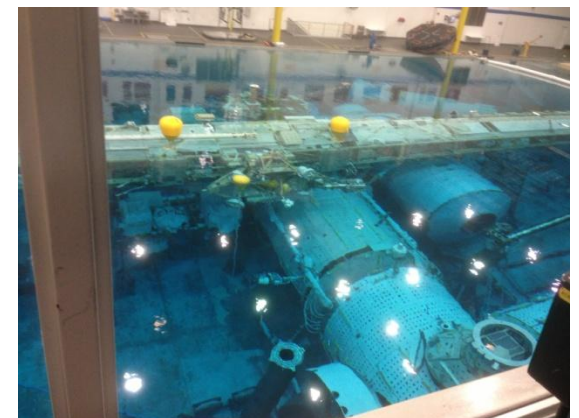
First experiments with monkeys

Irradiation with a proton
medical beam, 170 MeV



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

17th Meeting of the US/Russian Joint Working Group on Space Biomedical and Biological Sciences Research Houston, **NASA** (6-9 June 2015)

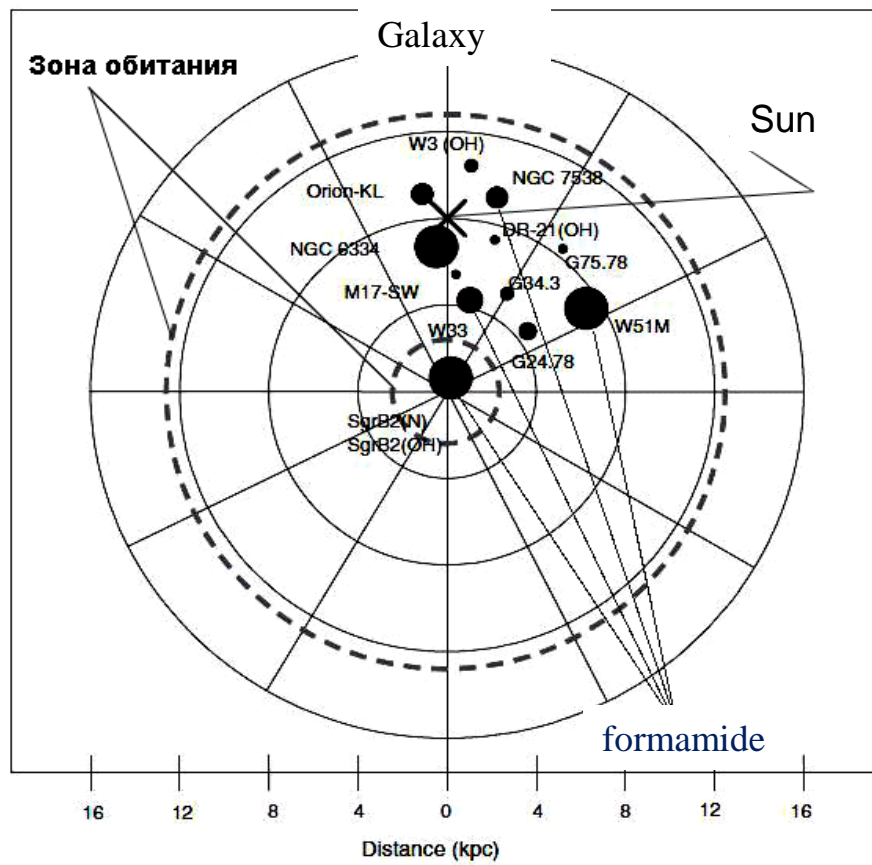
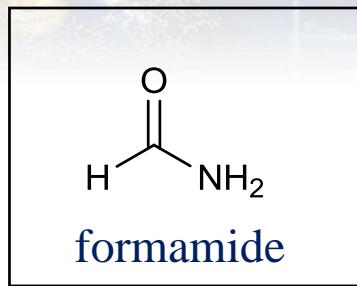
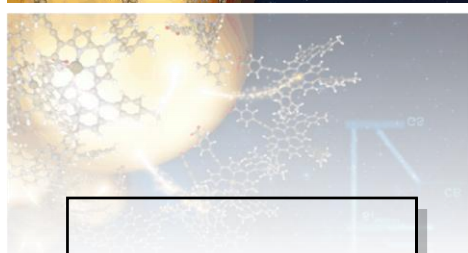


Astrobiology



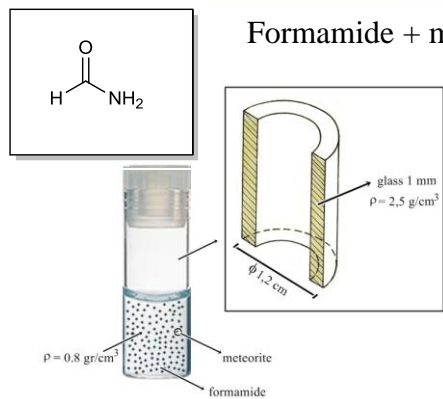
Gas-dust cloud Sagittarius B2, where formamide was found. Photo courtesy NASA

A model of the abundance and location of molecular clouds (the dotted line) containing formamide (black circles) in our Galaxy

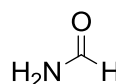
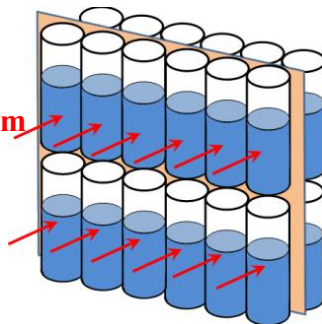


Astrobiology

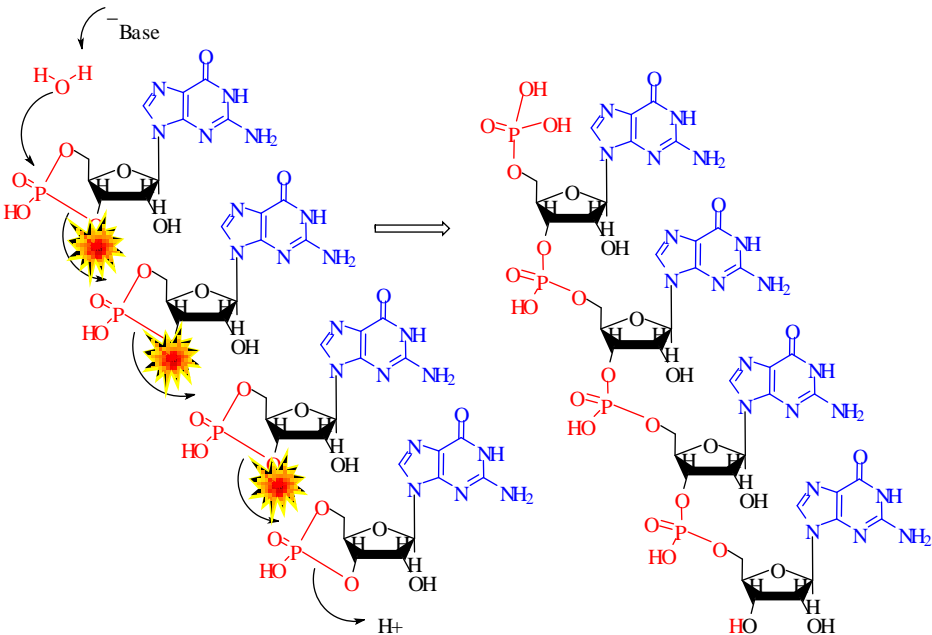
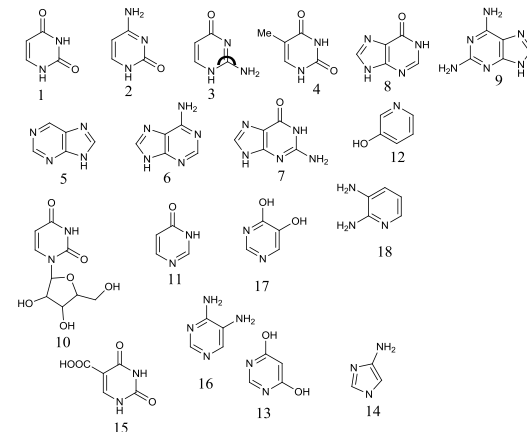
Collaboration: University of Viterbo, Sapienza University of Rome (Italy), and LRB



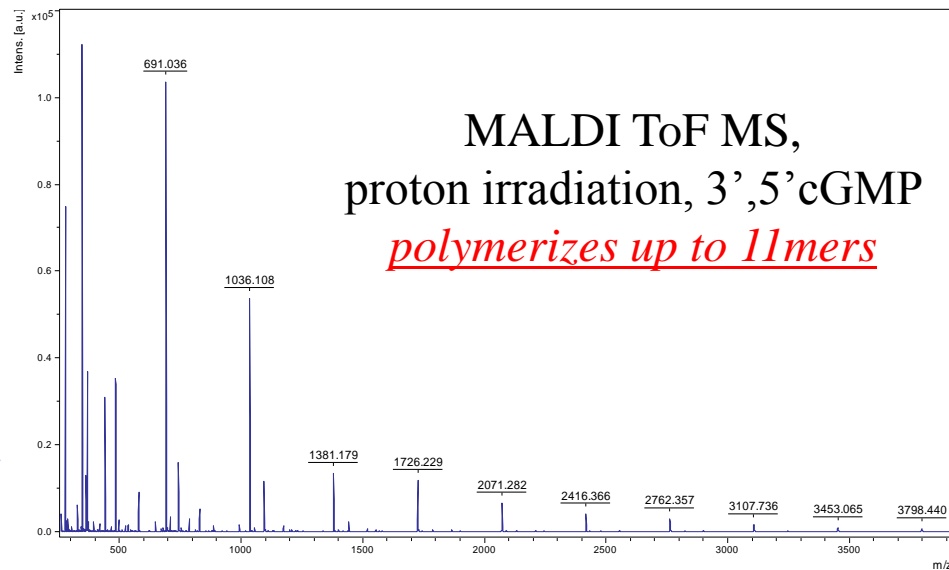
170 MeV protons
LET= 0.57 keV/ μm
t= 3min
T= 25°C



Radiation



m/z=1425



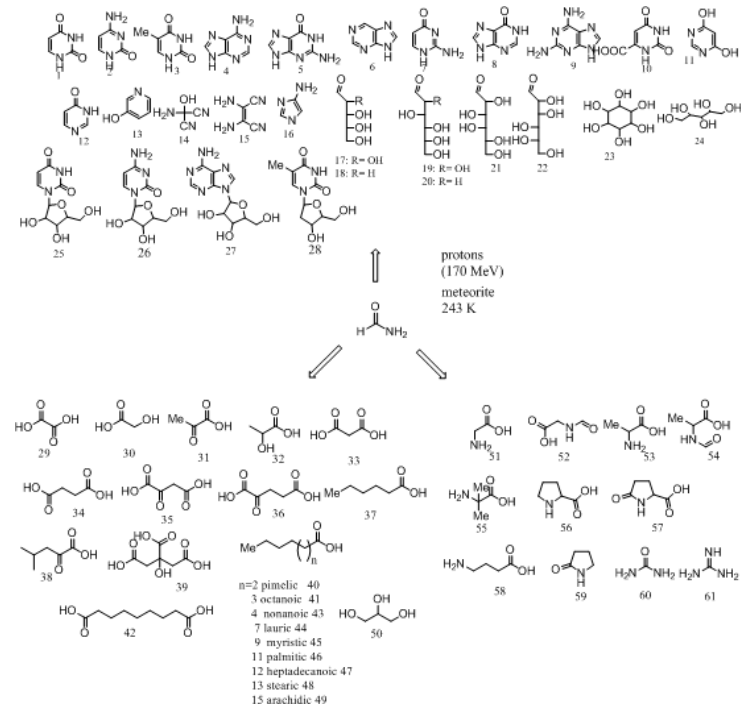
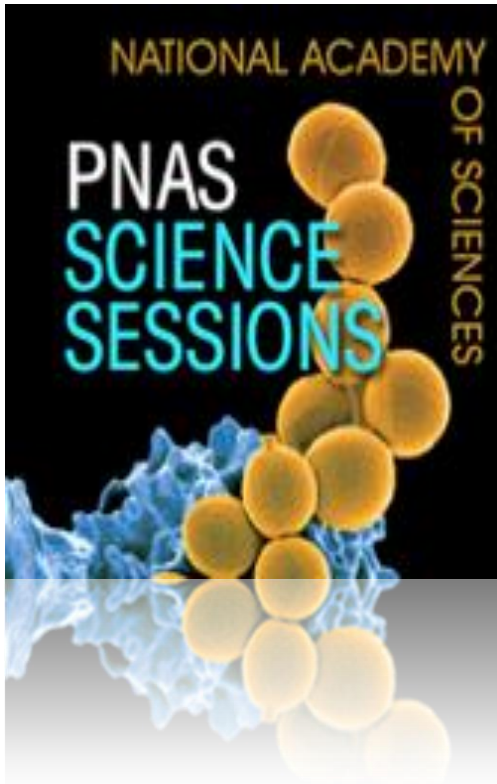
Astrobiology



The study was published in PNAS on 14.03.2015

Meteorite-catalyzed syntheses of nucleosides and of other prebiotic compounds from formamide under proton irradiation

Raffaele Saladino, Eleonora Carota, Giorgia Botta, Michail Kapralov, Gennady N. Timoshenko, Alexei Rozanov, Eugene Krasavin, and Ernesto Di Mauro. PNAS 10.1073/pnas.1422225112, 1-10



JINR's Accelerators for Radiobiology

Phasotron: protons 660 MeV



U-400M: heavy ions 50 MeV/u



Nuclotron: heavy ions up to 4 GeV/u



Thank you for your attention!