





WRMISS: Workshops on Radiation Monitoring for the International Space Station

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Agenzia Spaziale Italiana



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AstroParticle Experiments 4 Space Radiobiology

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In the near future, all the space agencies are working to restart the human exploration of space outside the Low Earth Orbit (LEO). Crewed space missions in this and the next decade will see the presence of humans on the Moon and Mars surface. One of the main showstoppers to be investigated for safe exploration and colonization is the biological effects of ionizing radiation that can compromise the health of astronauts/space workers.

The Astroparticle experiments presently operating in space (e.g., AMS02, ACE-Explorer, ...) could play a principal role in this vital task. Such experiments are actual cosmic ray observatories and a source of information crucial to investigating the fundamental physics open problems (e.g., Dark Matter, Antimatter) and improving the knowledge of radiobiology effects in space.

In this paper, a review of the past, present, and planned Astroparticle experiments operating would be presented and highlighted some of the possible contributions and improvements in the space radiobiology research field.

Also, will be presented some examples of progress in understanding the biological effects of radiation in space using the pieces of information acquired for astronomy and Astroparticle science and where such information has been used to enhance the space radiation field characterization and, consequently, improve crucial radiobiological issues in space (e.g., dose-effect models).

Finally, the use of the vast amounts of data taken from such experiments will open a new era of studies performed in different exposure scenarios that will allow a safe human space exploration outside of the Low Earth Orbit by addressing important radiation protection open questions, such as the dose relationship for cancer and non-cancer risk, the possible existence of dose threshold(s) for different biological systems and endpoints, and the possible role of radiation quality in triggering the biological response.

Outline

- AMS INFN Roma Sapienza Research Group
- Space Radiation Environment
- Cosmic ray Detectors in Space
- Possible sinergies
- ♦ Case Study :
 - ♦ Dose Effects Relationship (DER)
 - ♦ Target Effects vs Non Target Effects

INTRODUCTION

Alpha Magnetic Spectrometer (AMS) INFN ROMA SAPIENZA RESEARCH GROUP





SERVIZIO SANITARIO REGIONALE EMILIA-ROMAGNA Azienda Ospedaliero - Universitaria di Bologna



ALMA MATER STUDIORUM Università di Bologna

The AMS SPRB collaboration was created in 2017 by the synergy of the AMS INFN Roma Sapienza (Italy) group leaded by Alessandro Bartoloni with the medical physics research group leaded by Lidia Strigari currently at IRCCS university Hospital of Bologna (Italy)

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Alpha Magnetic Spectrometer AMS02

AMS is a particle detector measuring Galactic Cosmic Ray fluxes. It was installed on the International Space Station (ISS) on May 19, 2011







The AMS collaboration (http://ams02.space)



An international collaboration made of 44 Institutes from America, Asia and Europe

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The AMS02 detector has collected so far more than <u>200 billion</u> Cosmic Rays events.

More Info in the AMS-02 webpage:

https://amso2.space



Part of the AMS02 experiment was built at Rome (INFN & Sapienza)



The INFN Roma and the Sapienza university joined the AMS collaboration in 2001.

The group has taken part to the construction of the Transition Radiation Detector (TRD), having as main task the responsibility to develop the slow control electronics of the GAS System of the TRD (UG-Crate).

The UG-CRATE was part of a safety-critical system and the group took care of all the phases of the development (Design—Test-Integrate-Fly) following the NASA requirements.







Strategies for preventing radiobiological effects in space

Galactic Cosmic Rays induced Target and Non Target Effects in space

AMS02 Charged Particle characterization for Space Radiobiology investigations INFN Roma AMS-o2 wiki: https://wiki.infn.it/strutture/roma1/experiments/ams2/home

At INFN Roma AMS group, led by **Alessandro Bartoloni**, the primary activity is the use of the AMS measurements of cosmic rays to improve the space radiobiology knowledge with a primary emphasis on *the space radiation relevance and risk for human space exploration*.

In this topic, there is a strong collaboration and participation to the Roma group of the Medical Physics department of the IRCCS University Hospital of Bologna, led by Lidia Strigari.



Dr. Lidia Strigari

Medical Physics Department

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SPACE RADIATION & ASTRONAUT SAFETY

«To fully understand the relationship between ionizing radiation and biology, and to solve problems in this field, researchers incorporate fundamentals of **biology**, **physics**, **astrophysics**, **planetary science**, **and engineering**» (credit : NASA)

(credit : ESA)



(credit : CERN)

Cosmic Rays Interactions with the geo-magnetosphere

Earth is a cocoon !!!

Magnetosphere stops/deflects 99.9% of charged particles

the Earth Atmosphere is equivalent to a metal shielding 1 meter thick

The annual cosmic ray "dose" at sea level is around 0.27 mSv

> <10% of "background radiation" (Radon, Soils, Foods, Medical,..)



Origin of Space Radiation and Consequent Risk



From Strigari et Al Front. Public Health, 08 November 2021 | https://doi.org/10.3389/fpubh.2021.733337

Space Radiation composition

- Galactic Cosmic Rays (GCR)
- Particle emitted by the Sun (SEP) during isolated events
- Particle trapped in Earth's magnetic field (Radiation Belt)

None^{of} the 3 components is constant in time, mainly due to the solar activity

Human Space activities must cope with the high radiation environment of outer space.

Limits and concerns

The manned spaceflight especially the one beyond the LEO could represent a concern for the health of astronauts.

The limit in carrying out the missions are due to health effects

- short-term (<hours)</pre>
- acute effects (<months)</p>
- late effects including severe toxicity

Radioprotection in space is a difficult jobs due to the presence of different species of particle and nuclei that present different characteristics in penetrating the barrier and shielding

LEO-ISS X150-200 Moon **X300-400** we will go to the moon we S 300 kilometers from x250 (x750) Mars **Images Courtesy of NASA**

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A new era in human space exploration is coming ...



«Global Exploration Roadmap Lunar Surface Exploration Scenario update August 2020»

International Space Exploration Coordination Group (ISECG)

Figure 1. Updated ISECG Lunar Surface Exploration Scenario. A. Bartoloni - 25th WRMISS



The International Space Exploration Coordination Group (ISECG) is a forum set up by 14 space agencies to advance the Global Exploration Strategy through coordination of their mutual efforts in space explored and the set of the set

BLEO Space Exploration is restarted ! (IAC2021.11/2021)



Cosmic Ray Detectors in Space

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Energetic particles and completely ionized nuclei from outer space

Many orders of magnitude in energy and flux

E < 100 TeV: direct detection *E* > 100 TeV: detection of extensive-air-shower

The all-particle spectrum is a "power law" in many orders of magnitude of energy and intensity

with several features (*knee, ankle, ...*) $\gamma = 2,7$ for energy <100 TeV $\gamma = 3,3$ for energy >100 TeV

Credit C.Sparvoli

Timeline of Direct Measurement of CRs from 2000



3 possible platform for instruments and detectors baloons , Satellites , International Space Station

Cosmic Ray Observatory

"A **cosmic-ray** observatory is a scientific installation built to detect high-energyparticles coming from space called **cosmic rays**.

This typically includes photons (high-energy light), electrons, protons, and some heavier nuclei, as well as antimatter particles.

Cosmic Rays in the Heliosphere Synchrotron bremss.IC lower the IS CR

Credit : P. Picozza (INFN)

PAMELA

Principal Operating Cosmic Ray Space Detectors

Satellite Based









Cosmic Ray Components Identification

e+,e-	📀 ALL
p+,p-	ALL
D,He	🖉 ALL
Low-Z (<=8)	ALL (PAMELA up to Z=6)
Middle-Z	AMS02, CALET, ISS-CREAM, ACE, DAMPE
High-Z (>14)	MISO2, CALET, ISS-CREAM, ACE, DAMPE



Properties of Iron Primary Cosmic Rays: Results from the Alpha Magnetic Spectrometer AMS Collaboration • M. Aguilar (Madrid, CIEMAT) et al. (Jan 29, 2021)

Published in: Phys.Rev.Lett. 126 (2021) 4, 041104

Properties of Heavy Secondary Fluorine Cosmic Rays: Results from the Alpha Magnetic Spectrometer

AMS Collaboration • M. Aguilar (Madrid, CIEMAT) et al. (Feb 25, 2021)

Published in: Phys.Rev.Lett. 126 (2021) 8, 081102

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ANSO2 PUBLISHED CRS A. BARTOLONI - 25TH WRMISS

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Cosmic Rays Solar Modulation

Cosmic rays from interstellar medium are «screened» by the Heliosphere.

This effect is particularly visible at low energies

Measurements of time evolution of cosmic ray fluxes of different particles over an extended period of time is very valuable Properties of Daily Helium Fluxes

AMS Collaboration • M. Aguilar (Madrid, CIEMAT) et al. (Jun 10, 2022) Published in: *Phys.Rev.Lett.* 128 (2022) 23, 231102 Periodicities in the Daily Proton Fluxes from 2011 to 2019 Measured by the Alpha Magnetic Spectrometer on the International Space Station from 1 to 100 GV

AMS Collaboration • M. Aguilar (Madrid, CIEMAT) et al. (Dec 27, 2021)





FIG. 1. The daily AMS proton fluxes for six typical rigidity bins from 1.00 to 10.10 GV measured from May 20, 2011 to October 29, 2019 which includes a major portion of solar cycle 24 (from December 2008 to December 2019). The AMS data

Short term Solar Modulation of GCR Daily Proton and Helium Fluxes and Helium to Proton flux ratio

Properties of Daily Helium Fluxes

AMS Collaboration • M. Aguilar (Madrid, CIEMAT) et al. (Jun 10, 2022) Published in: *Phys.Rev.Lett.* 128 (2022) 23, 231102 Periodicities in the Daily Proton Fluxes from 2011 to 2019 Measured by the Alpha Magnetic Spectrometer on the International Space Station from 1 to 100 GV

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Short term Solar Modulation of GCR Daily Proton and Helium Fluxes and Helium to Proton flux ratio

Cosmic Rays in the Magnetosphere



(Figure credits: NASA)

Particles trapped in the Earth magnetic field create regions of high radiation called Van Allen belts. The ISS crosses one of the belts over South America, causing a sudden increase of the observed radiation known as the South Atlantic Anomaly.

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«Ten Years of AMS» Session

The South Atlantic Anomaly²^{as} Seen by AMS

Incoming particle rate at the poles and in the SAA is high. This causes low collection efficiency, mostly in the inner part of the SAA. However, the efficiency is high on the external sides of the SAA.



M. Aguilar et al., Phys. Rep. 894 (2021) 1–116.

Energetic particle with charge up to 2 are known to exist in this region. While there is no previous observation of energetic (R>1GV) Z>2 particles inside SAA.

Credit A.Oliva & AMS Collaboration

Stably Trapped Nuclei in the SAA

Backtracing allows to select particles stably-trapped in Earth's magnetosphere. A clear population of stably trapped ions (Z>2) entering in AMS both from the **top** and the **bottom** has been identified.



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- 10 years of AMS-02 data have been used to look for ions with Z>2 below geomagnetic cutoff;
- A stably trapped population has been clearly identified below 5 GV in the SAA region;
- This population has properties (rigidity, charge, arrival direction) distinctly different from GCRs;
- This is a high-Z, high-energy population (up to 5 GV) never observed before.

Credit A.Oliva & AMS Collaboration

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GCR sensitivity analysis

- Identifications of CR components of the CR that are of interest for the computation of possible risks associated with the manned exploratory space missions in LEO and BLEO scenarios.
- Use of space radiation sensitivity studies we also recognised that they correspond with the data taken by the astroparticle experiments

- ♦ Environmental GCR model : BON2010^[4] ♦ ICRP 60 Radiation Quality Factors ♦ ICRP 103 for Tissue Weights
- "FAX": Female Adult voXel phantom^[5] ٢
- ♦ Transport Code : HZETRN- π /EM^[6]





1) Environmental Model Characterization:

- Use the enormous data at energies > 1 GeV
- Improve affects the accuracy and precision of the risk assessment potentially underestimating the actual damage.

2) Dose Equivalent Estimation:

- Measurements only of absorbed doses, by passive dosimeters, are insufficient for investigating biological effects or assessing radiation risk for astronauts.
- LET distributions, their QFs (up to 30), and RBE of high-LET particles constituting the space radiation environment.

3) Transport Code Validation :

- Monte Carlo (MC) simulation code can be further implemented to better describe the interaction with the matter of GCR environments thanks to the improvement of accuracy of cross sections at high energy of elementary particles (electrons, protons), light and heavy nuclei (Helium to Iron and beyond).
- The implementation of transport code at these energies allows predicting the particle interactions with the known geometries of installed detectors. The determination of ray / particle tracing, energy spectrum and deposited energies collected in several materials can serve for a subsequent MC transport code validation (e.g. through a possible Bayesian approach).
- The calculations of dose equivalents allow generating an accurate and precise database for subsequent MC simulation codes validation applied to human tissues. Moreover, MC codes can be used for designing ad hoc shielding of spacecrafts and space landers.

4) Space Exposition Scenario Dose Computation:

- Implementation of Montecarlo codes to calculate the dose and so predict/describe the effects of GCR particles interacting with cells, tissues/organs and astronauts, which can be modeled as geometries with increasing details and complexities.

5) Ground or Space based Experiment setup definition:



A synergy with Astroparticle researches

A Case Study :

Dose Effects Relationship - Target Effects vs Non Target Effects

Dose-Effect Relationship

Crucial point is to predict the toxicity of the space radiation expected for the astronauts/space workers and the creation of reliable *mathematical models* that describe the correlation between the exposition to IR and the possible damages to the organs at risk

Aim: to implement a platform including the more reliable dose-effect models for space radiation, we developed an ad hoc software in R-script language



Target Effects (TE) vs Non Target Effects (NTE)

- Non-targeted effects (NTEs) include bystander effects where cells traversed by heavy ions transmit oncogenic signals to nearby cells, and genomic instability in the cells progeny.
- Studies on the Harderian gland, chromosomal aberrations at low dose and many mechanistic studies support the NTE model, with evidence of a **supra-linear effect at low doses** of NTE compared to a linear effects for TE
- This NTE are expected also at the dose-rates that occur in space.

Non-Targeted Effects Models Predict Significantly Higher Mars Mission Cancer Risk than Targeted Effects Models

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Materials & Methods: Hazard Function for Tumor Prevalence (TP)

Prevalence is the number of people/cell with a specific disease or condition in a given population at a specific time. This measure includes both newly diagnosed and pre-existing cases of the disease.

Tumor prevalence (TP) is described by a Hazard function, H, which is dependent on radiation type for γ -rays while for charged particles is dependent on the charge number (Z), kinetic energy (E) and fluence (F).

$$TP = 1 - e^{-H(\Sigma,\Sigma,\Gamma)}$$
$$H_{\gamma} = H_0 + \left[\alpha_{\gamma} D + \beta_{\gamma} D^2 \right] * S(D)$$
$$H_{CP}(Z, E, F) = H_0 + \left[\Sigma F + \beta D^2 \right] * S(D)$$

-H(7 E E)

Where:

- ➢ H0 represents the background prevalence
- > α_{γ} and β_{γ} are the linear and quadratic coefficient with dose Induction terms
- > Σ is pseudo-biological action cross section taking in account the particle track structure models
- ➤ S(D) is the *Cell Survival Probability*.

Results: R-script Library includes the most used Cells Survival Probability models

- 1. Theory n-target N-hit model (nTNH) Two special case of nTNH including:
 - Theory single Target single hit model (sTSH)
 - Theory single Target N-hit model (sTNH)
- 2. Theory Linear Quadratic Model (LQ)
- 3. Linear Quadratic Model modified by hyperradiosensitivity(HRS) effect.
- 4. Theory Linear Quadratic Cubic Model (LQC) for high dose.
- Sublesion Theory Repair misRepair Model (S-RMR)
- 6. Sublesion Theory Lethal potentially lethal Model (S-LPL)
- Sublesion Theory Saturable Repair Model (S-SR)

1.
$$S(D) = 1 - (1 - B)^n, B = e^{\frac{-D}{D_0}} [1 + \sum_2^N \frac{\binom{D}{D_0}^{N-1}}{(N-1)!}]$$

2. $S(D) = e^{-\alpha D - \beta D^2}$
3. $S(D) = exp\{-\alpha \left(1 + \left(\frac{\alpha_s}{\alpha} - 1\right)e^{\frac{-D}{D_0}}\right)D - \beta D^2\}$
4. $S(D) = e^{-\alpha D - \beta D^2 - \gamma D^3}$
5. $S(D) = e^{-\alpha D} [1 + \left(\frac{aD(1 - e^{(-\lambda T)})}{\varepsilon}\right)]^{\varepsilon \phi}$
6. $S(D) = e^{-(n_L - n_{PL})D} [1 + \frac{n_{PL}D}{\varepsilon} (1 - e^{-\varepsilon_{PL}t_r})]^{\varepsilon}$
7. $S(D) = e^{-\frac{n_0 - C_0}{1 - \frac{C_0}{n_0}e^{kT(C_0 - n_0)}}}$

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Materials and Methods: Experimental Data Set (Alpen et. al. 1997)

Prevalence of Harderian Gland Tumors

- Gammas 55.5TBq Co60
- Hydrogen with energy 250A, LET 0.4 KeV/ μ m
- Exposition time in between 60 sec. to 120 sec.
- Irradiation field is 3 x 5 cm².
- Background Prevalence is H0 = 0.026

TABLE VIIPrevalence of Harderian Gland Tumors after Irradiationwith Protons, Neon Ions, and Niobium Ions

		<u>, ,,,,,</u>				
lon and energy	Dose (Gy)	Number	With tumors	At risk	Prevalence ⁴ (%)	
Controls	0	198	155	4	2.6 ± 2.5	
Protons (250A MeV)	0.40 0.80 1.60 3.20	47 42 48 28	43 41 43 7	44 8 13 24	9.3 ± 6.1 19.5 ± 12.1 30.2 ± 13.7 29.2 ± 18.2	

TABLE II
Prevalence of Harderian Gland Tumors
after ⁶⁰ Co Gamma Irradiation

		Mice				
Dose – (Gy) N	Number	At risk	With tumors	Prevalence ^a (%)		
0	198	155	4	2.6 ± 2.5		
0.4	292	229	11	4.8 ± 2.7		
0.8	278	161	15	9.3 ± 4.5		
1.6	244	117	16	13.7 ± 6.2		
3.2	181	115	37	32.2 ± 8.5		
7.0	90	52	24	46.2 ± 13.6		

" ±95% CI.



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Hazard Function Targe Effect (TE) vs Non-Target Effects (NTE)

The NTE model assumes a nonlinear type response in addition to the linear dose term at low doses.

The η function represents the NTE contribution, which is parameterized as a function of the particle Linear Energy Transfer (L).

We tuned the radiobiological parameters to reproduce available experimental data $H_{TE}(Z, E, F) = H_0 + \left[\Sigma F + \beta D^2\right] * S$

 $H_{NTE}(Z, E, F) = \left[H_0 + \Sigma F + \beta D^2 + \eta\right] * S$

$$\eta = \eta_0 L e^{-\eta_1 L} [1 - e^{-N_B ys}]$$

Where:

>

- L is the Linear Energy Transfer of the particle
- \succ $N_{Bys} = F$ luence $* A_{Bys}$
 - A_{Bys} is the number of bystander cells surrounding a cell traversed directly from a HZE particle that receive an oncogenic signal.

Results: Effective Pseudo-Biological Action Cross Section $\Sigma(Z,E) = \Sigma_0 P(Z,E) + \frac{\alpha_{\gamma}L}{6.24} [1 - P(Z,E)], P(Z,E) = [1 - e^{\left(-\frac{Z^{*2}}{kv_c^2}\right)}]^m$ $\Sigma(1,250) = \Sigma_0 P(Z,E) + \frac{0.55*0.4}{6.24} [1 - P(1,250)], P(1,250) = [1 - e^{\left(-\frac{Z^{*2}}{kv_c^2}\right)}]^3$

Where:

- * Σ_0 and k are parameters of his cellular track structure model
- * α_{γ} is the linear regression coefficient for acute doses of γ -rays for the same endpoint
- * Z^* is the effective charge number of the particle,
- Vc is the particle velocity relative to the velocity of light
- ★ m is the number of target in a single cell



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Results: α_H and β_H Calculation

 $TP_{H} = 1 - e^{-H_{H}(1,250,F)}$

 $H_{TE}(1, 250, F) = 0.026 + [\Sigma(1, 250) * F + 0.1092 * D^2] * S$

 $H_{NTE}(1, 250, F) = 0.026 + \left[\Sigma(1, 250) * F + 0.1092 * D^2 + \eta_H\right] * S$

 $\eta_H = 0.00048 * 0.4e^{-0.00281 * 0.4} \left[1 - e^{-216 * F}\right]$

Where:

* β_H is the quadratic coefficient with dose Induction terms, irradiation for hydrogen

For the cell survival probability is used the target theory n-target N-hit model (nTNH) with n = 3, N = 1

Using the experimental data of 250A MeV Hydrogen irradiation we calculate the $\alpha_{\rm H}$, $\beta_{\rm H}$ induction term



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Results: TE vs NTE for Protons

Calculation of the TE and NTE TP models showing for Proton 250A Mev there is no relevant differences in the tumor Prevalence versus dose as expected.





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Results: Tool for NTE components evaluation



Calculation of the NTE with different models and for different exposition scenarios

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Summary

- In the coming years there will be a great interest for space human mission non only to explore but also for a permanent presence of humans in LEO and outside the geo-magnetosphere
- Space Radiation is a main concern and the first one showstopper in many human exploration scenarios.
- Astroparticle Experiments are a principal source of information to perform this investigations complementary to what is usually done in the research field.
- Dose-Effects models knowledge could benefit from the use of such information, a synergy with the experience from the clinical field is crucial to perform this task
- At INFN ROMA AMS Group we are developing a platform including the more reliable dose-effect models for space radiation and the AMS data

«To fully understand the relationship between ionizing radiation and biology, and to solve problems in this field, researchers incorporate fundamentals of **biology, physics, astrophysics, planetary science, and engineering.»** (credit : NASA)



THANKS FOR THE ATTENTION !

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AMS02 INFN ROMA and Sapienza University Web Site

BACKUP SLIDE

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Dose-Effects Relationship

The known dose-effect relationships are based on a limited number of astronauts

(hundreds)

Total Space Radiation Dose (mGy)	<0.2	0.2-1.99	2-3.99	4-10.99	≥11	Total
# Astronauts	14	19	11	15	14	73
# Cancer Deaths	2	2	1	0	2	7
# Cardiovascular Disease Deaths	1	4	1	1	0	7
# Accident Deaths	6	5	0	0	1	12
# Other Deaths	1	0	1	0	1	3
# Unknown Deaths	1	0	0	3	1	5
Mean Medical Dose (SD)	2.4 (6.4)	27.7 (13.6)	34.4 (20.8)	29.1 (15.6)	32.5 (21.7)	25.1 (19.4)
Mean Year at Birth (SD)	1932.6 (4.1)	1931.7 (5.2)	1931.6 (2.5)	1932.2 (4.4)	1931.5 (3.3)	1931.9 (4.1)
Mean Age at Entry into Astronaut Corps (SD)	31.6 (2.7)	32.2 (3.4)	33.0 (2.5)	31.8 (2.8)	32.5 (2.2)	32.2 (2.8)
Mean Follow up Time (SD)	29.3 (23.6)	40.3 (15.0)	46.4 (12.9)	50.7 (7.8)	48.1 (7.5)	42.8 (16.1)
Total Group Person Years	409.9	766.5	510.1	760.8	673.4	3120.8
Mean Age at Death (SD)	57.7 (23.8)	65.7 (15.9)	64.5 (14.9)	78.2 (19.9)	74.9 (10.2)	65.2 (19.1)
Mean Current Age of Living Astronauts (SD)	79.9 (2.9)	82.1 (3.9)	84.9 (3.1)	83.6 (3.6)	83.8 (2.3)	83.4 (3.4)

Table 1. Early astronaut cohort demographics binned by total space radiation dose category. SD = standard deviation.

Radiation Exposure and Mortality from Cardiovascular Disease and Cancer in Early NASA Astronauts S.Robin et Al - 2018





Needs of an improvements

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REVIEW article

Front. Public Health, 08 November 2021 Sec.Radiation and Health https://doi.org/10.3389/fpubh.2021. 733337 This article is part of the Research Topic Medical Application and Radiobiology Research of Particle Radiation View all 16 Articles >

Dose-Effects Models for Space Radiobiology: An Overview on Dose-Effect Relationships

Lidia Strigari¹,

Silvia Strolin¹, Alessio Giuseppe Morganti² and

We did and publish in 2021 an extensive review of the existent literature



Dose-Effects Models for Space Radiobiology:

An Overview on Dose-Effect Relationships

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Space radiobiology is an interdisciplinary science that examines the biological effects of ionizing radiation on humans involved in aerospace missions. The dose-effect models are one of the relevant topics of space radiobiology. Their knowledge is crucial for optimizing radioprotection strategies, the risk assessment of the health hazard related to human space exploration, and reducing damages induced to astronauts from galactic cosmic radiation. Dose-effect relationships describe the observed damages to normal tissues or cancer induction during and after space missions.

Based on a *PubMed* search including 53 papers reporting the collected **dose-effect relationships after space missions or in ground simulations**, 7 significant dose-effect relationships (e.g., eye flashes, cataract, central nervous systems, cardiovascular disease, cancer, chromosomal aberrations, and biomarkers) have been identified.

For each considered effect, the absorbed dose thresholds and the uncertainties/limitations of the developed relationships are summarized and discussed. The current knowledge on this topic can benefit from further *in vitro* and *in vitro* radiobiological studies, an accurate characterization of the quality of space radiation, and the numerous experimental dose-effects data derived from the experience in the clinical use of ionizing radiation for diagnostic or treatments with doses like those foreseen for the future space missions.

The growing number of pooled studies could improve the prediction ability of dose-effect relationships for space exposure and reduce their uncertainty level. Novel research in the field is of paramount importance to reduce damages to astronauts from cosmic radiation before Beyond Low Earth Orbit exploration in the next future. The study aims at providing an overview of the published dose-effect relationships and illustrates novel perspectives to inspire futureresearch.

Model	Study Type	Dose Range/Threshold or LET	#Papers	Reliability	Priority
Eye Flashes	Spaceflight	LET>5-10 KeV/µm	4	****	•
Cataract	Spaceflight	8 mSv	5	***	***
CNS	Ground/Simulations	100-200 mGy	11	**	*****
CVD	Spaceflight	1000 mGy	4	*	***
	Ground/Simulations	0.1-4,500 mSv	8		
Cancer	Spaceflight	< 100 mGy	2	***	*****
	Ground/Simulations	< 100 mGy	9		
Biomarkers or	Spaceflight	<5-150 mGy	11	***	*****
Chromosomal Aberrations	Ground /Simulations	< 10,000 mGy	4		
Other Risks	Ground/Simulations	2,000 mGy	2	*	***
1486 gf America 319 7 236 200		826			
1906 200 300 400	500 600 700	800 800	-		

We made and publish in 2021 an extensive review of the existent literature to use as starting point for improvements this research areas

https://doi.org/10.3389/fpubh.2021.733337

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REVIEW article

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Dose-Effects Models for Space Radiobiology: An Overview on Dose-Effect Relationships

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Alessandro Bartoloni^{3*}

Model	Study Type	Dose Range/Threshold or LET	#Papers	Reliability	Priority		
Eye Flashes	Spaceflight	LET>5-10 KeV/µm	4	****	*		
Cataract	Spaceflight	8 mSv	5	***	***		
CNS	Ground/Simulations	100-200 mGy	11	**	****		
CVD	Spaceflight	1000 mGy	4	*	***		
	Ground/Simulations	0.1-4,500 mSv	8				
Cancer	Spaceflight	< 100 mGy	2	***	****		
	Ground/Simulations	< 100 mGy	9				
Biomarkers or Chromosomal Aberrations	Spaceflight	<5-150 mGy	11	***	****		
	Ground /Simulations	< 10,000 mGy	4				
ol Other Risks	Ground/Simulations	2,000 mGy	2	*	***		
*= Very Low, **=Low, ***=Medium, **** = High, **** = Very High.							

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THE RESEARCH TOPIC INITIATIVE

Research Topics are Open Access themed article collections (similar in nature to classical special issues) with:

a dedicated landing page, Continuous publication, Advanced impact metrics, Cross-disciplinarity, Multiple article types, e-book production



Astroparticle Experiments to Improve the Biological Risk Assessment of **Exposure to Ionizing Radiation in the Exploratory Space Missions**

The actual and next decade will be characterized by an exponential increase in the exploration of the Beyond Low Earth Orbit space (BLEO). Moreover, the firsts tentative to create structures that will enable a permanent human presence in the BLEO are forecast. In this context, a detailed space radiation field characterization will be crucial to optimize radioprotection strategies (e.g., spaceship and lunar space stations shielding, Moon / Mars village design, ...), to assess the risk of the health hazard related to human space exploration and to reduce the damages potentially induced to astronauts from galactic cosmic radiation. On the other side, since the beginning of the century, many astroparticle experiments aimed at investigating the unknown universe components (i.e., dark matter, antimatter, dark energy, ...) have been collecting enormous amounts of data regarding the cosmic rays (CR) components of the radiation in space.

Such experiments essentially are actual cosmic ray observatories, and the collected data (cosmic ray events) cover a significant period and permit to have integrated not only information of CR fluxes but also their variations on time daily. Further, the energy range is exciting since the detectors operate using instruments that allow measuring CR in a very high energy range, usually starting from the MeV scale up to the TeV, not usually covered by other space radiometric instruments. Last is the possibility of acquiring knowledge in the full range of the CR components and their radiation quality. The collected data contains valuable information that can enhance the space radiation field characterization and, consequently, improve the radiobiology issues concerning one of the most relevant topics of space radiobiology represented by the dose-effect models

This articles collection accepts original research papers and review papers relating (but not limited to) the following topics:

- The analysis and proposal on how to use these astroparticle experiments data to enhance the space radiation field characterization and, consequently, improve the radiobiology issues in space concerning one of the most relevant topics of space radiobiology represented by the dose-effect models and relationship

- The proposal of new methods or instruments to use the astroparticle experiments to improve the space radiobiology knowledge (i.e., real-time dosimetry, monitoring of solar activities, ...)

Keywords: Cosmic Ray, Space Radiation, Space Radiobiology, Astro-Particle Experiments, Human Space Exploration

Participating Journals

Manuscripts can be submitted to this Research Topic via the following journals:

Frontiers in Astronomy and Space Sciences Astrobiology

Frontiers in Physics **Radiation Detectors and Imaging**

Frontiers in Public Health

Radiation and Health



frontiers in Astronomy and Space Sciences

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A new scientific language is needed to support the exploratory space missions because of the return of humans outside the Low Earth Orbit. The keywords are Peacefully, Safely, Transparently.

In that context, a priority is to keep the space exploration community secure and safe, and a crucial part is a detailed and accurate ionizing radiation health effects characterization.

Participate in creating part of this new language oining this interdisciplinary Frontiers Research Topic!



Improve the Radiation Health Risk Assessment for Humans in Space Missions

Since 2018, the INFN Roma Sapienza AMS group has collaborate with researchers and scientists to investigate the possibilities of using the CRD to improve the radiation health risk assessment for humans in space missions.

Collaborations were mainly focused on creating synergy within different scientific communities (radiobiology, medical physics, radiotherapy, and nuclear medicine) and institutions (Research, Universities, Space Agencies and Industry).

In 2019 we organize at INFN Roma Sapienza a thematic meeting with participants from ESA and Thales Alenia Space

SPACE RADIOBIOLOGY AND PRECISION GALACTIC COSMIC RAY MEASUREMENTS

ON HOW THE AMSO2 EXPERIMENT ON THE INTERNATIONAL SPACE STATION CAN HELP THE RADIATION HEALTH HAZARD ASSESSMENT IN EXPLORATORY SPACE MISSIONS

> Lunedì 4 novembre 2019 Dipartimento di Fisica – Aula Conversi



14:30-14:45 Introduzione A. Bartoloni – INFN Roma



14:45-15:35 High precision measurements of charged cosmic rays in space with the Alpha Magnetic Spectromete M. Paniccia, Università di Ginevra



15:35-16:20 ESA Human Spaceflight Radiation Research Programme activities. L. Surdo, European Space Agency

16:20-17:05 Shielding design for long duration human exploratory space missions : issues and future perspective. *M. Girauda, Thales Alenia Space*



The Research Topic Initiative

- While progressing in the research activity raised the awareness that to make progress in such a field it was required a new scientific language able to connect and create **synergy** between different scientific communities.

 Firstly, cause to understand the relationship between ionizing radiation and biology and to solve problems in this field, researchers incorporate fundamentals of biology, physics, astrophysics, planetary science, and engineering.

- Further **space exploration and colonization** collects the worldwide hopes of a new era characterized by transparency and peacefully development. In that sense, these expectations coincide with the primary scientific interest, and science could play a breakthrough role in such direction.

- Among the many possibilities thus, we decided, supported and asked by the Frontiers Editorial team, to launch this research topic named "Astroparticle Experiments to Improve the Biological Risk Assessment of Exposure to Ionizing Radiation in the Exploratory Space Missions".

TOPIC EDITORS







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Gianluca Cavoto



We created a research topic **editorial board** that was representative of **different scientific cultures** and **geographic areas** and invited many researchers and scientists from many different research areas due to the strong interdisciplinarity of the topic.



Space Weather

RESEARCH ARTICLE

10.1029/2020SW002456

Key Points:

- The Badhwar-O'Neill 2020 GCR model is presented
- The updated model is calibrated using new AMS-02 and PAMELA data
- Solar activity is described using ACE/CRIS daily integral flux measurements

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The Badhwar-O'Neill 2020 GCR Model

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Abstract The Badhwar-O'Neill (BON) model has been used for some time to describe the galactic cosmic ray (GCR) environment encountered in deep space by astronauts and sensitive electronics. The most recent version of the model, BON2014, was calibrated to available measurements to reduce model errors for particles and energies of significance to astronaut exposure. Although subsequent studies showed the model to be reasonably accurate for such applications, modifications to the sunspot number (SSN) classification system and a large number of new high-precision measurements suggested the need to develop an improved and more capable model. In this work, the BON2020 model is described. The new model relies on daily integral flux from the Advanced Composition Explorer Cosmic Ray Isotope Spectrometer (ACE/CRIS) to describe solar activity. For time periods not covered by ACE/CRIS, the updated international SSN database is used. Parameters in the new model are calibrated to available data, which include the new Alpha Magnetic Spectrometer (AMS-02) and Payload for Antimatter Matter Exploration and Light-nuclei Astrophysics (PAMELA) high-precision measurements. It is found that the BON2020 model is a significant improvement over BON2014. Systematic bias associated with BON2014 has been removed. The average relative error of the BON2020 model compared to all available measurements is found to be <1%, and BON2020 is found to be within $\pm 15\%$ of a large fraction of the available measurements (26,269 of 27,646 \rightarrow 95%).