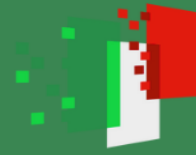




Ministero
dell'Università
e della Ricerca



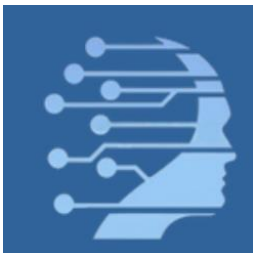
PNC
Piano nazionale per gli investimenti
complementari al PNRR
Ministero dell'Università e della Ricerca



Geant4 Monte Carlo simulations for medical physics applications and novel radiotherapy techniques

Giuliana Milluzzo

Istituto Nazionale di Fisica Nucleare Sezione di Catania



Computing@CSN5: applications and innovations at INFN
Bari, 14-16 Ottobre 2024

giuliana.milluzzo@ct.infn.it



A bit of history...

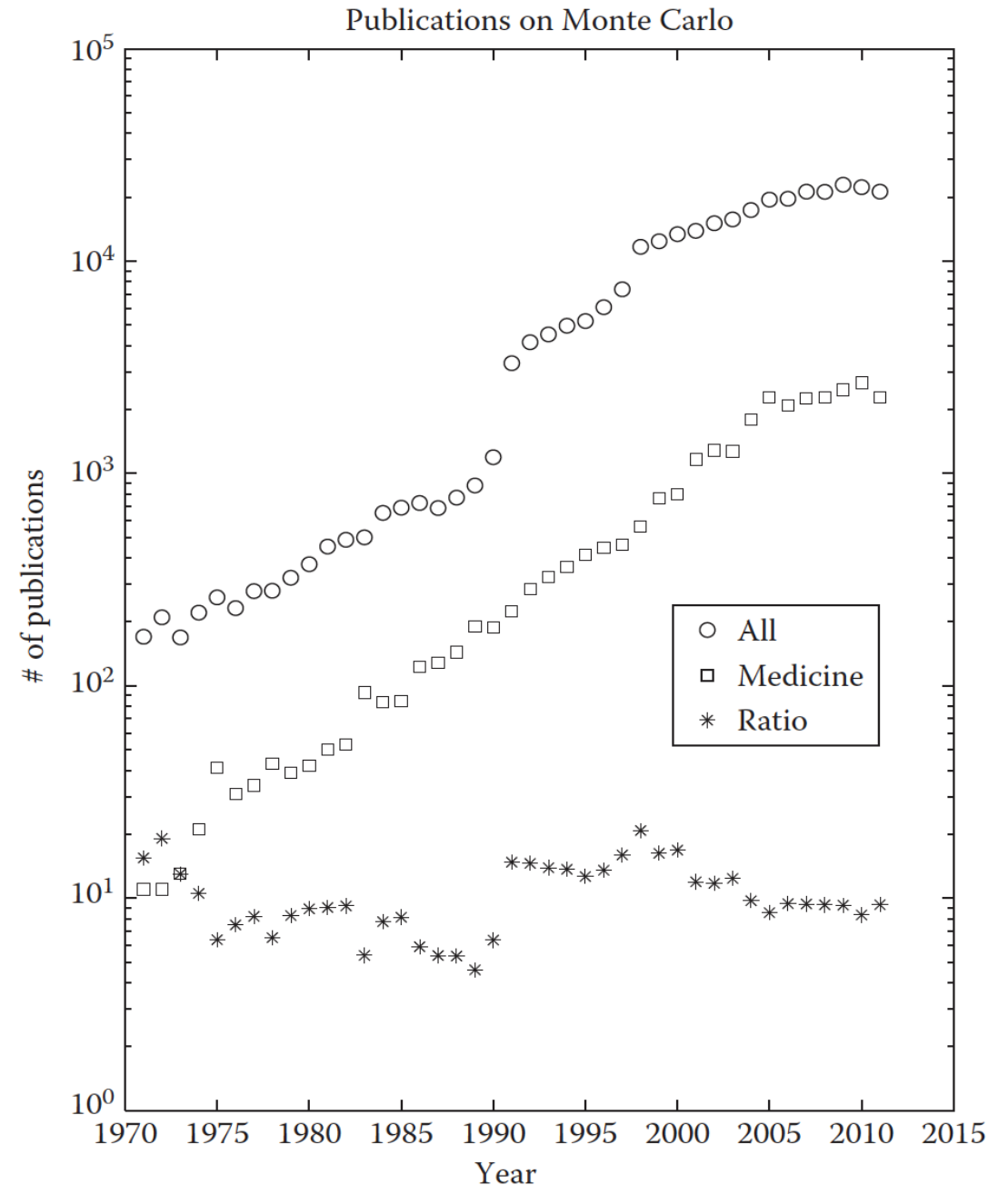
In **1949**: First paper on the Monte Carlo methods.
(**"The Monte Carlo Method**, Metropolis and Ulam)

1950-52: *First papers employing the Monte Carlo method using electron transport for Medical Physics*, by Robert R. Wilson

1954: The **Symposium on Monte Carlo Methods** with 70 attendees, many of whom would be recognized as "founding fathers" by Monte Carlo practitioners in the radiological sciences.

1958-60: first use of an **electronic digital computer** in simulating high-energy cascades by **Monte Carlo methods for Medical Physics** reported by Butcher and Messel (independently by Varfolomeev and Svetlollobov).

Publications using Monte Carlo methods **rapidly increased**
10% of them related to **medical applications**



Monte Carlo simulations for medical physics applications

Predicting physical quantities of interest in medical physics

Dose, energy, fluence distributions, beam spot size for radiotherapy, diagnostics, TPS

Computing physical/biological quantities difficult to measure

LET, RBE, microdosimetric spectra

Modelling, design of new devices, beamlines in conventional-RT and for innovative radiotherapy modalities

Choose a reliable tool for the simulation of these processes

Validation of physical models and continuous benchmarking with experimental data

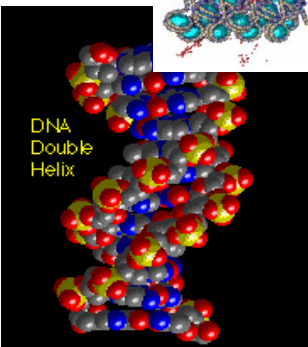
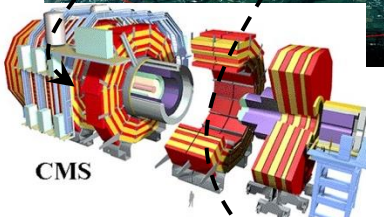
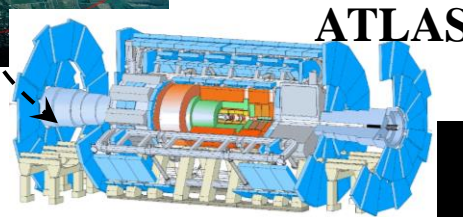
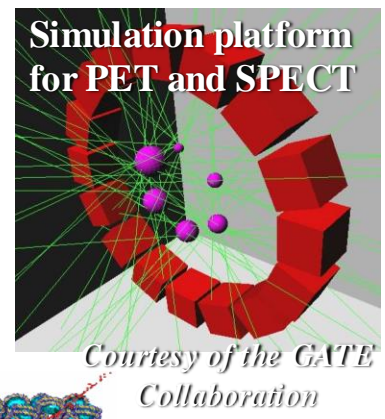
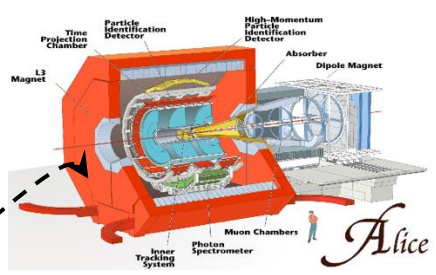
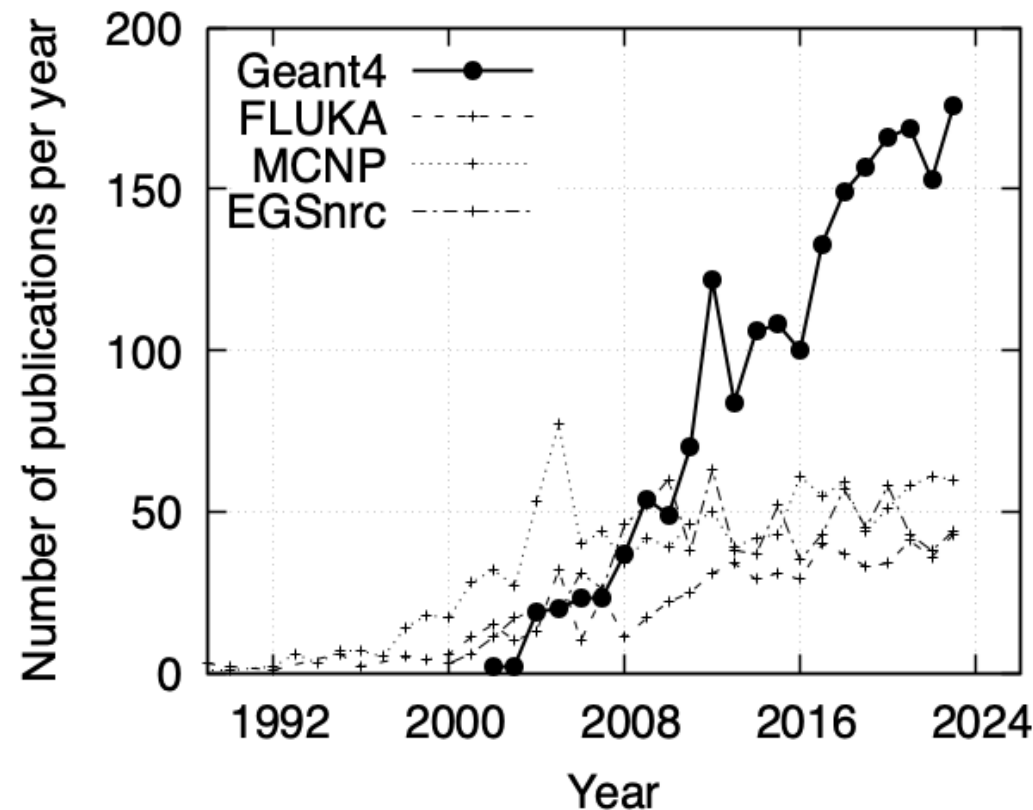


The Geant4 Monte Carlo toolkit

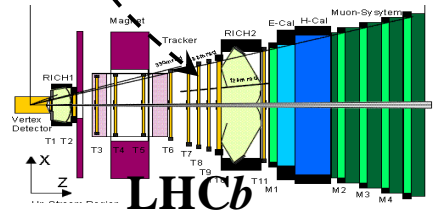


- 1974:** First version of the code released at CERN for high energy particles
- 1983:** Geant3 written in Fortran language
- 1998:** Geant4 written in C++ object oriented language was released for the simulation of large scale HEP experiments at CERN (Geneva)

Geant4 Medical applications



Courtesy of S. Incerti, Bordeaux



The Geant4 Medical Simulation Benchmarking Group (G4MED)

<https://twiki.cern.ch/twiki/bin/view/Geant4/G4MSBG>

P Arce¹, J Archer², **L Arsini³**, D Bolst², J M C Brown⁴, B Caccia⁵, A Chacon⁶, **P Cirrone⁷**, M A Cortés-Giraldo⁸, D Cutajar², **G Cuttone⁷**, P Dondero⁹, A Dotti¹⁰, B Faddegon¹¹, **S Fattori⁷**, C Fedon¹², S Guatelli^{2,*}, A Haga¹³, S Incerti¹⁴, V Ivanchenko¹⁵, D Konstantinov¹⁶, I Kyriakou¹⁷, A Le², Z Li¹⁴, M Maire¹⁸, A Malaroda¹⁹, **C Mancini-Terracciano³**, A Mantero⁹, C Michelet¹⁴, **G. Milluzzo²⁰**, **F Nicolanti³**, M Novak²¹, C Omachi²², **L Pandola⁷**, J. H. Pensavalle²³, A Perales²⁴, Y Perrot²⁵, **G Petringa⁷**, S Pozzi⁵, J M Quesada⁸, J Ramos-Méndez¹¹, **F Romano²⁰**, A B Rosenfeld², M Safavi-Naeini⁶, D Sakata²⁶, L G Sarmiento²⁷, T Sasaki²⁸, Y Sato¹³, **A Sciuto⁷**, I Sechopoulos^{29,30}, E C Simpson³¹, R. Stanzani⁹, A. Tomal³², T Toshito²², N H Tran¹⁴, C. White², D H Wright¹⁰. ***INFN**

- Created in 2014.
- **Coordination Team since 2018:**
 - **Coordinator:** Susanna Guatelli (Univ. Wollongong, Australia)
 - **Deputy-coordinator:** Pedro Arce (CIEMAT, Spain)
- 57 researchers; 32 institutions from 12 different countries.



MEDICAL PHYSICS
The International Journal of Medical Physics Research and Practice

Research Article | [Full Access](#)

Report on G4-Med, a Geant4 benchmarking system for medical physics applications developed by the Geant4 Medical Simulation Benchmarking Group

P. Arce, D. Bolst, M.-C. Bordage, J. M. C. Brown, P. Cirrone, M. A. Cortés-Giraldo, D. Cutajar, G. Cuttone, L. Desorgher, P. Dondero, A. Dotti, B. Faddegon, C. Fedon, S. Guatelli, S. Incerti ... See all authors

First published: 11 May 2020 | <https://doi.org/10.1002/mp.14226> | Citations: 102

+ Second paper under review in Medical Physics



Courtesy of S. Guatelli

G4-Med tests

Electromagnetic physics tests

- **Brachytherapy** (Ir-192 and I-125)
- Electron **FLASH radiotherapy**
- MV X-ray **radiotherapy test**
- Photon attenuation coeff
- Electron electronic stopping power
- Electron backscattering
- 13 MeV electron forward scattering
- **Bremsstrahlung** from thick targets
- Fano cavity

Geant4-DNA tests

- Low dose energy electron Dose Point Kernels
- Microdosimetry
- Chemistry

Hadronic nuclear cross section tests

- Nucleus-nucleus hadronic inelastic scattering cross sections
- **62 MeV/u ^{12}C fragmentation**
- Charge-changing cross sections for 300 MeV/u ^{12}C ions

Electromagnetic + hadronic physics tests

- **62 MeV proton** beam test (cell survival modelling and averaged LET track)
- In-vivo PET for carbon ion therapy
- 67.5 MeV proton Bragg curves in water
- Light Ion Bragg Peak curves
- Neutron yield of 113 MeV and 256 MeV protons and 290 MeV/u carbon ions
- **Fragmentation test** of a 400 MeV/u ^{12}C ion beam in water

Goals:

Provide physics list recommendations

Monitor physics capability of Geant4 through the different version of the code

Electromagnetic physics models

Opt3, **Opt4**, Livermore, Penelope, SS



Hadronic physics models

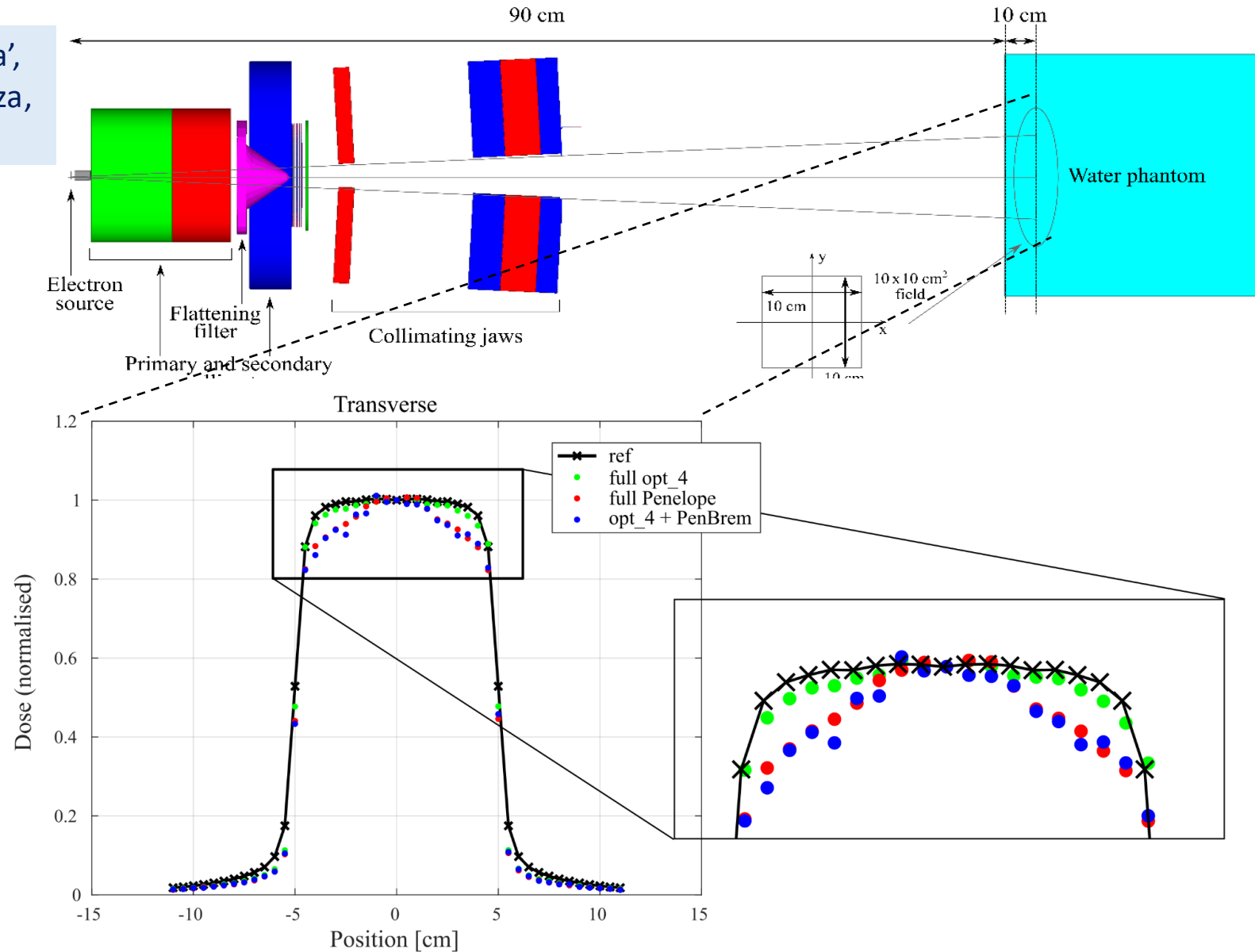
The **Binary Cascade Model** (BIC), Quantum Molecular Dynamics (QMD), Liege Intranuclear Cascade (INCL)

Conventional radiotherapy

The MV X-ray radiotherapy test

By B. Caccia and S. Pozzi (Istituto Superiore di Sanita', Rome, Italy), and C. Mancini-Terracciano (La Sapienza, Rome, INFN-RM1, Italy)

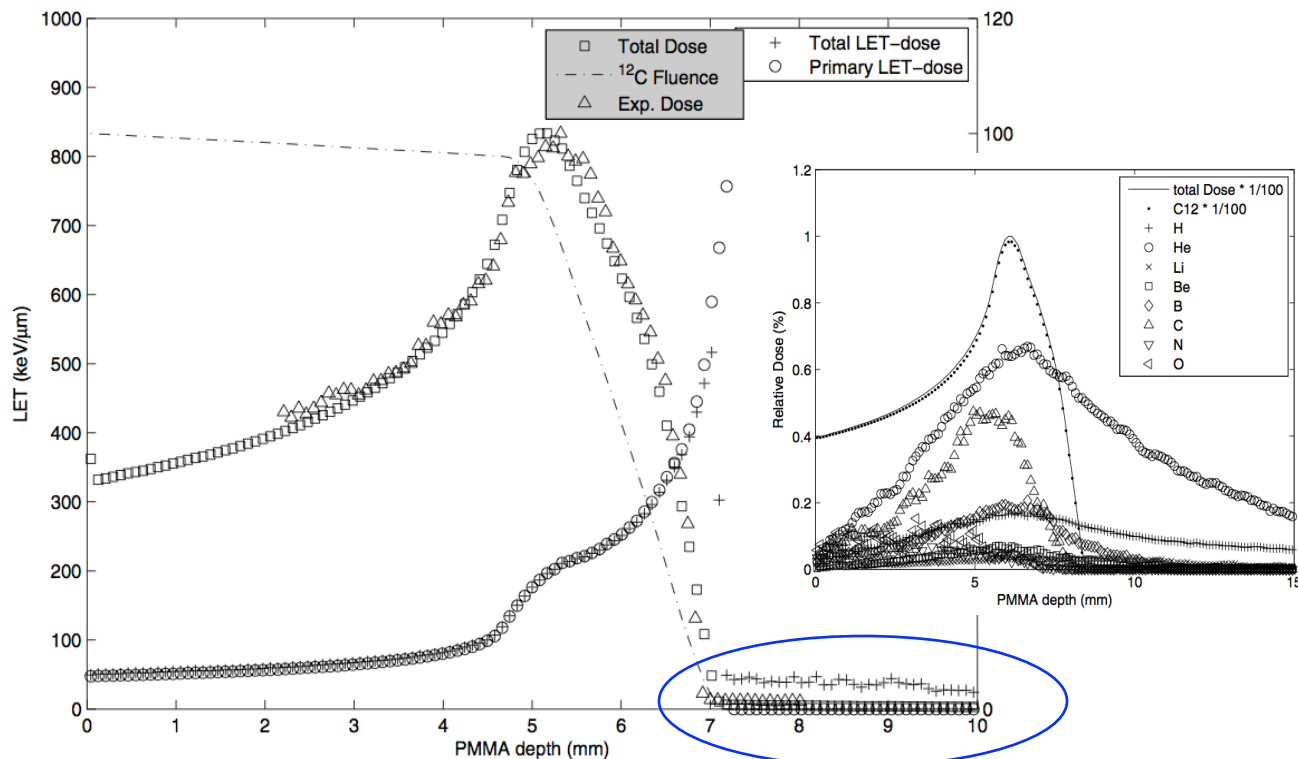
- Geant4 advanced example **medical_linac**
- Model of a GE Saturne 43 linear accelerator (EURADOS Report Caccia et al, 2020-05)
- The Geant4 Penelope Bremsstrahlung process seems to have problems
- This is now under investigation



Proton and hadrontherapy

^{12}C fragmentation tests

C. Mancini-Terracciano (Uni-Ro, INFN-RM1), S. Guatelli (CMRP), D. Bolst, C. Omachi, T. Toshito (Nagoya PTC), T. Sasaki (KEK)

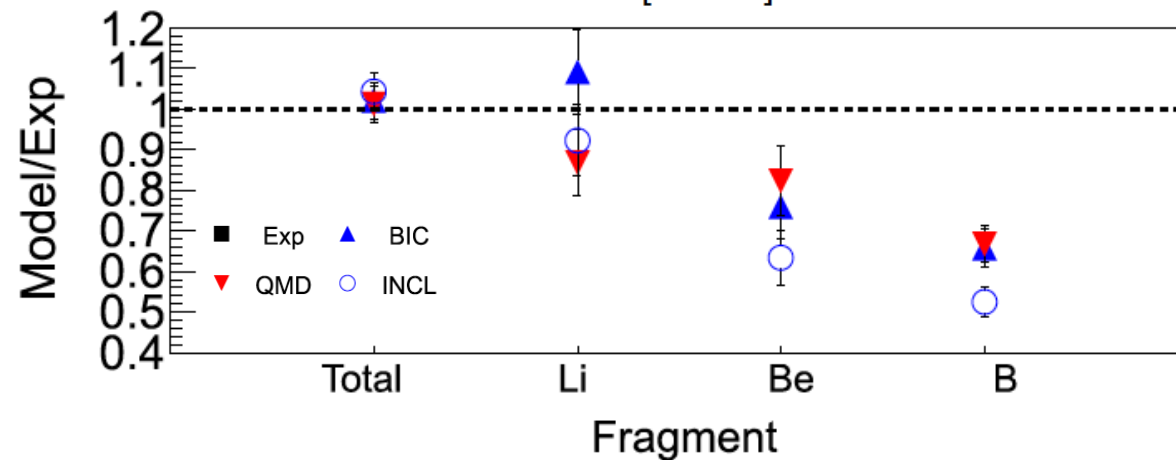
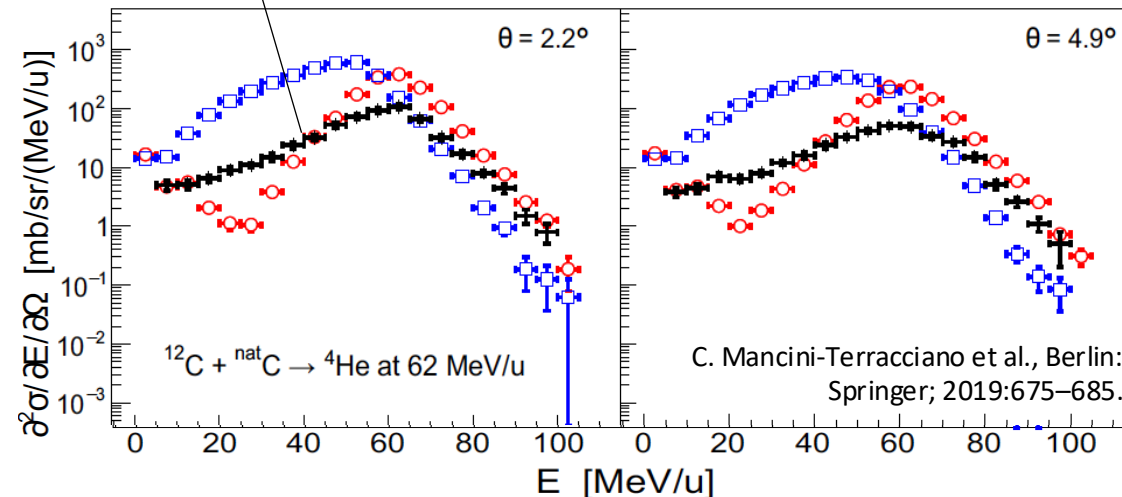


Romano *et al.* PMB 59 (2014)

Increased biological effects respect to the predictions

FRAGG & FRATT experiment@ INFN-LNS
De Napoli *et al.* PMB 59 (2014)

Alpha particle spectra



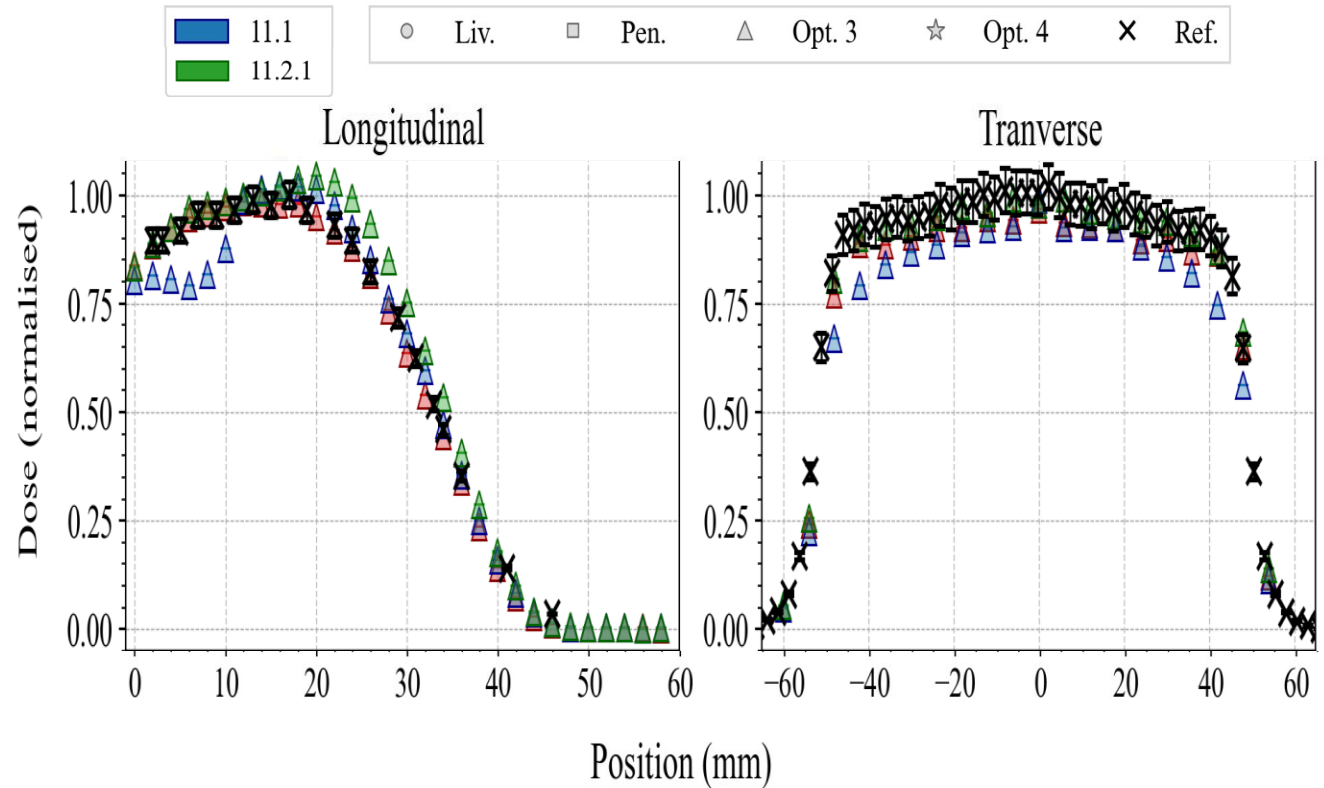
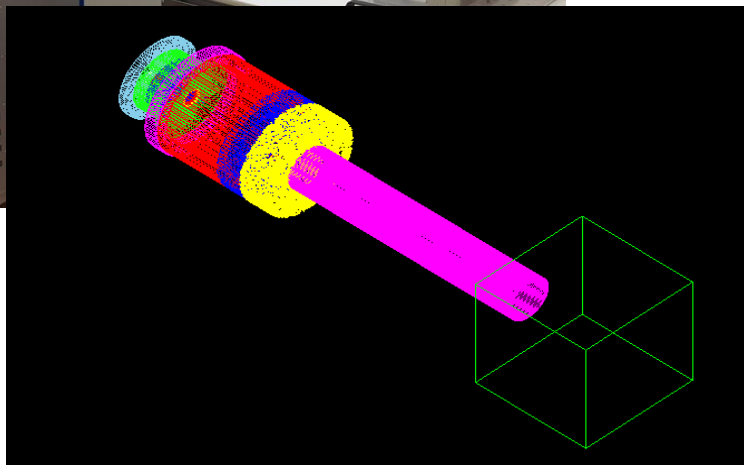
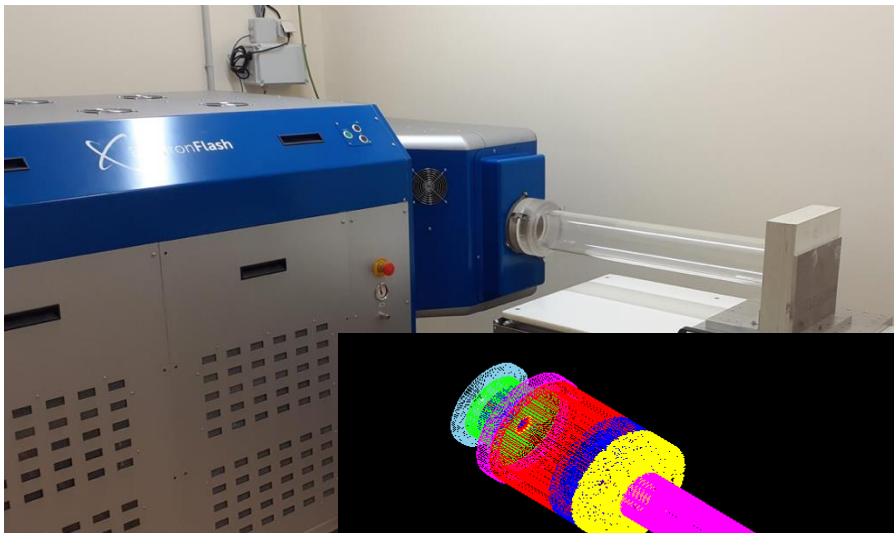
ARCE *et al.*, Med Phys, 48 (1), 2021

FLASH radiotherapy

Electron FLASH radiotherapy test

G. Milluzzo & F. Romano (INFN Catania), and J. H Pensavalle (Azienda Ospedaliero Universitaria Pisa)

Triode Electron Gun Equipped **ElectronFlash** Linac, manufactured by Sordina Iort Technologies S.p.A installed at the **CPFR (Centro Pisano Flash Radiotherapy)** in Pisa, Italy



eFLASH_radiotherapy impact within the national projects



FRIDA
CSN5



CSN5



DREAM
CNTT



Spoke4 (WP3)



Spoke8 (WP2)

Silicon carbide array Detector for dose profile measurements at FLASH regimes



R4 Research for Innovation

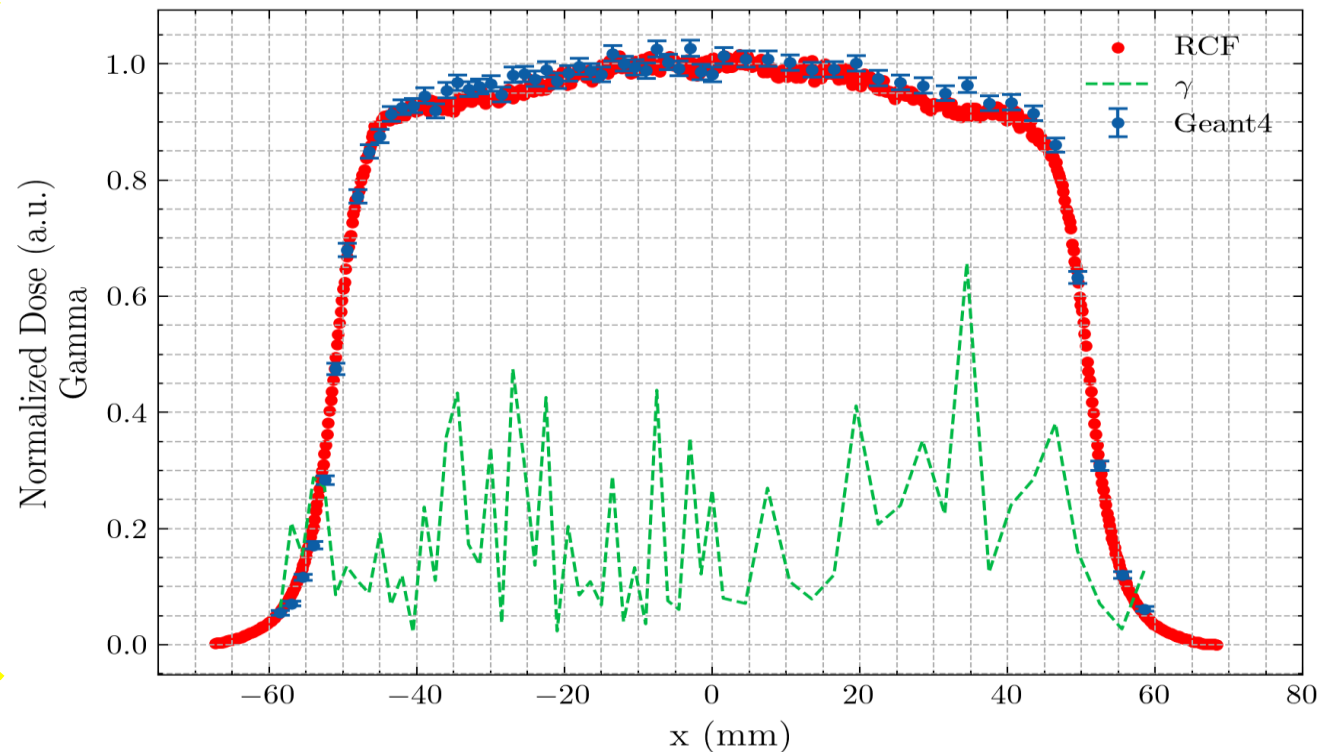
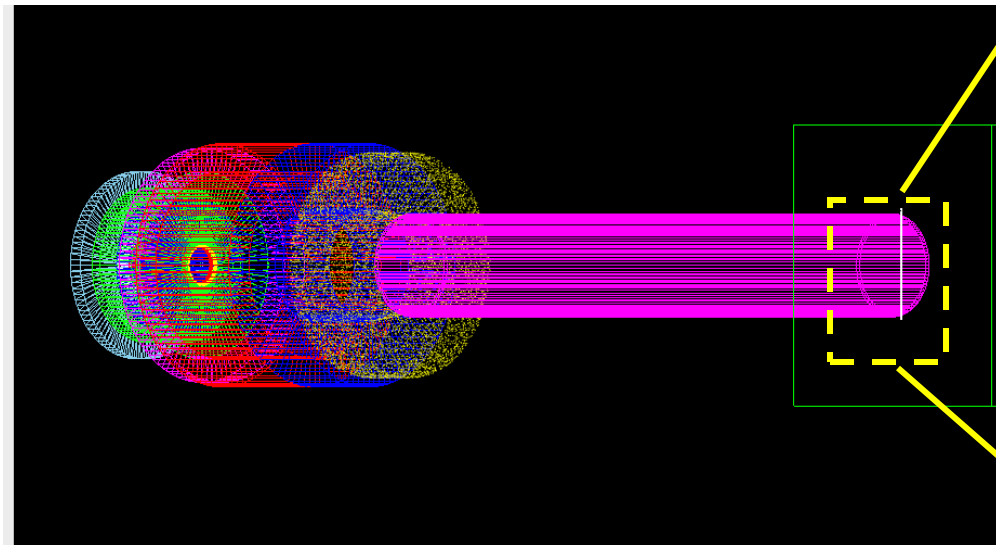
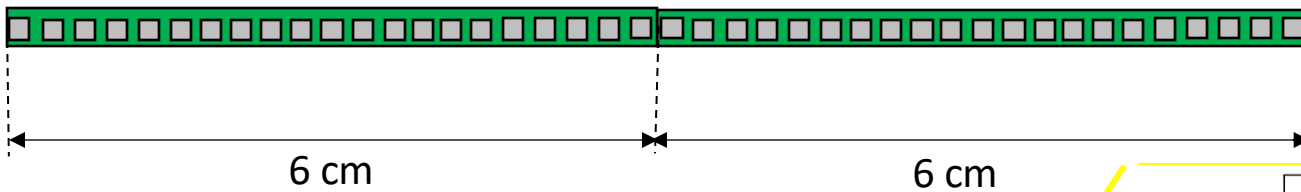


DREAM

1. Reducing the total amount of delivered dose
2. Keeping a high spatial resolution
3. Minimize the cost and the complexity → number of detectors

FLASH RADIOTHERAPY

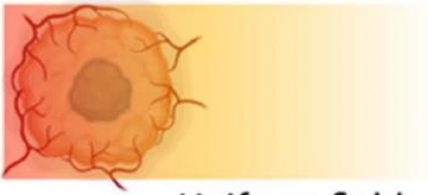
Dose per pulse: 1-20 Gy
Instantaneous Dose Rate: > 40 Gy/s
Irradiation Time: <200 ms



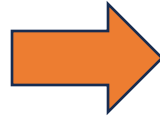
Applications in minibeam radiotherapy

PI: F. Di Martino (INFN-PI)
F. Romano (INFN-CT)

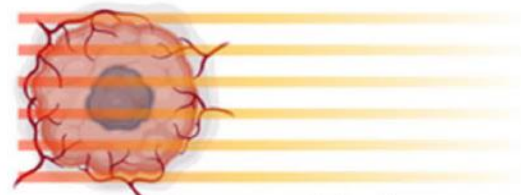
Conventional-RT



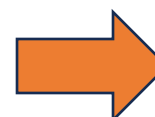
Uniform field



Minibeam-RT (MBRT)



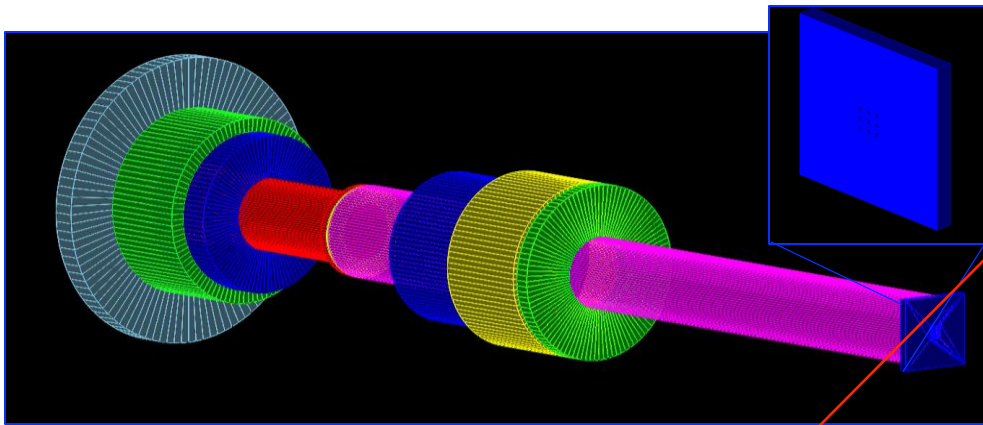
>100 μm wide beams



Minibeam facilities

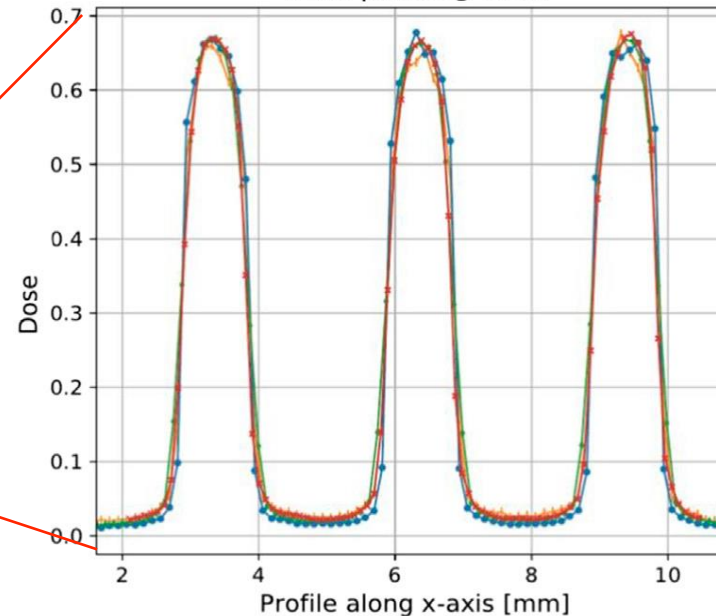
- **Electron** minibeam @ CPFR & INFN-TO & INFN-CT
- **Proton** minibeam @ Trento Protontherapy Centre

Minibeam collimator design for the ElectronFLASH linac in Pisa

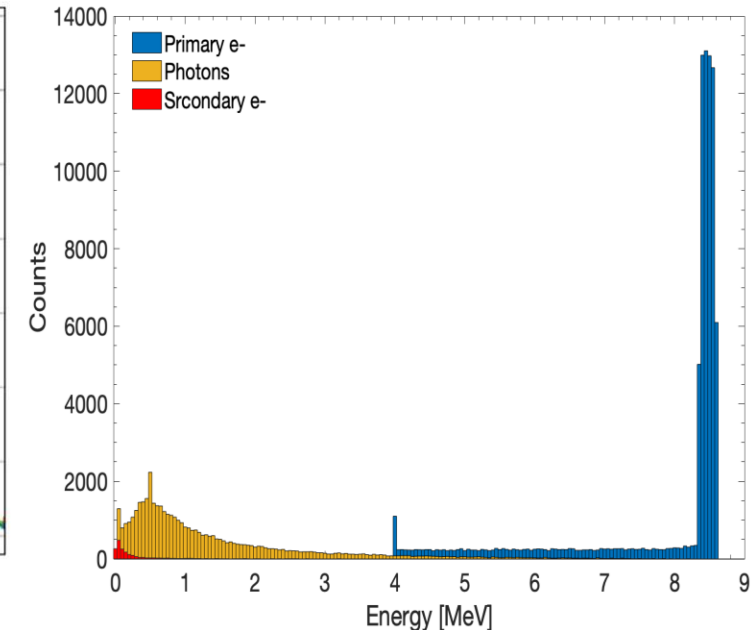


J. Pensavalle et al., Frontiers Physics 2023

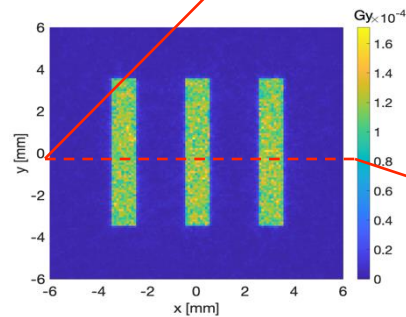
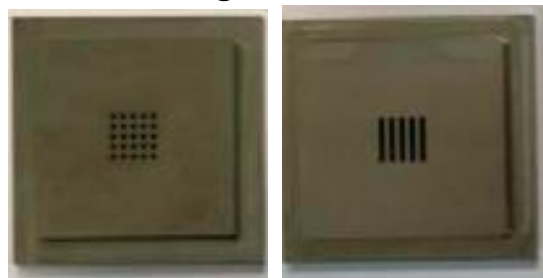
Dose profile @ 0mm



Simulations of secondary particles contamination



Real tungsten collimators



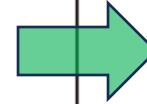
Evaluation of microdosimetric parameters

Microdosimetry: measurement of the statistical fluctuations of energy deposition in site size (cellular and sub-cellular scale) → biological effects as the result of energy deposition in sensitive sites or targets of specific size.

The *lineal energy*, y , is the quotient of ε_s by \bar{l} , where ε_s is the energy imparted to the matter in a given volume by a single energy-deposition event, and \bar{l} is the mean chord length of that volume, thus

$$y = \frac{\varepsilon_s}{\bar{l}}$$

Unit: J m^{-1}



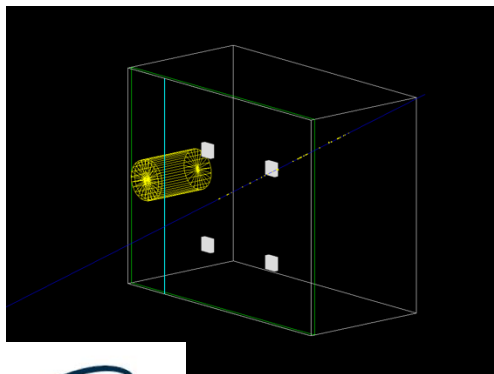
Measurement of microdosimetric spectra

TEPC-based detectors
Solid state detectors

Colautti P et al., *Front. Phys.* 8:550458.
V. Conte et al., *Phys. Med. Biol.* 65 (2020) 245018

Monte Carlo track structure and condensed history codes

exp_microdosimetry Geant4 advanced example

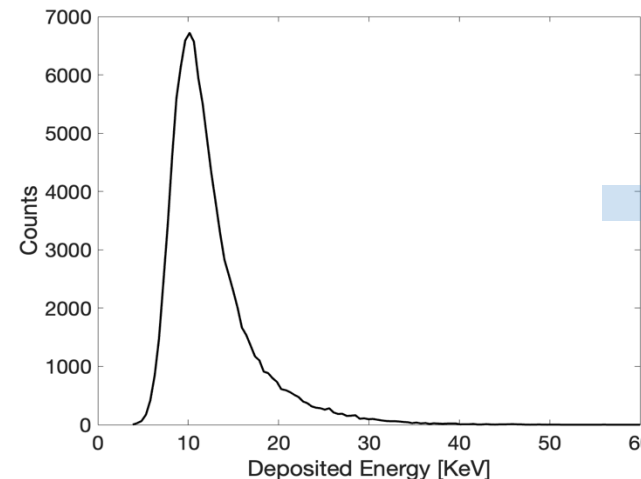


Microdosimeters

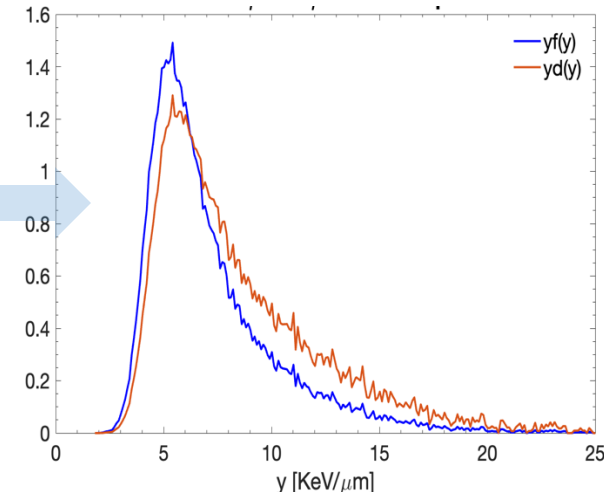
- Simplified diamond microdosimeter
- The microdiamond detector (TorVergata)
- Bridge silicon microdosimeters
- Diamond telescope
- **SiC microdosimeter**
- **Mini TEPC**



20 MeV protons,
SiC 100x100 $\mu\text{m}^2/2 \mu\text{m}$



Microdosimetric Kinetic Model (MKM)
$$S(D) = \exp - \left(\alpha_0 + \frac{\beta}{\rho \pi r_d^2} \bar{y}_D \right) D + \beta D^2$$

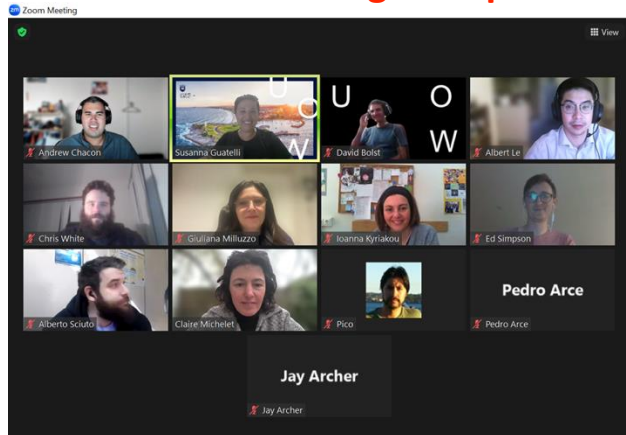


Conclusions and remarks

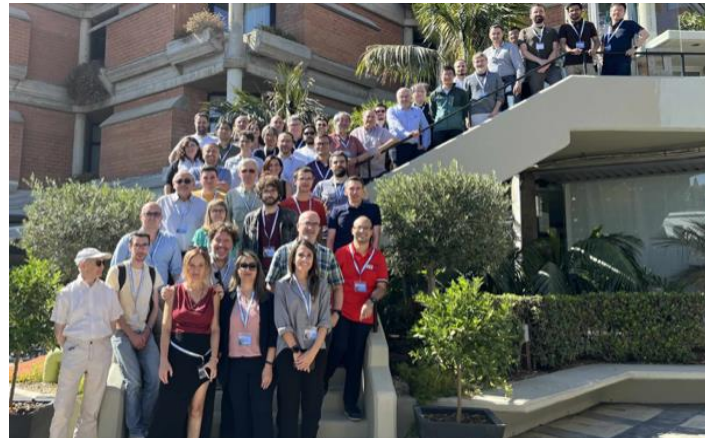
- **Geant4** is an open source toolkit maintained by the international collaboration continuously validated against experimental data
- The **G4-Med** project born to benchmark the physical models in the domain of medical physics application, has proved:
 - To monitor how changes in the Geant4 physics component translate in physical quantities of interest;
 - To supports significantly the development of the Geant4 physics component.
- **Geant4 applications** and simulations are actively used in **INFN** and **PNRR** projects

Acknowledgements

The Geant4 Medical Simulation Benchmarking Group



Geant4 International collaboration



INFN-CT Research group

Francesco Romano, Giuliana Milluzzo, Chinonso Okpuwe, Gabriele Trovato, Said Ahmed

CPFR

F. Di Martino, J. Pensavalle, L. Masturzo



DREAM

FRIDA

**Thank you for
your attention**

