

Master projet Hadronthérapie

AG du GdR Mi2B
Marseille, 29 Juin – 1 Juillet 2026

Responsables scientifiques :

Sara MARCATILI, Marie Vanstalle,
Claire Rodriguez-Lafrasse, Juliette Thariat
on behalf of Hadrontherapy IN2P3 project

Responsable technique :

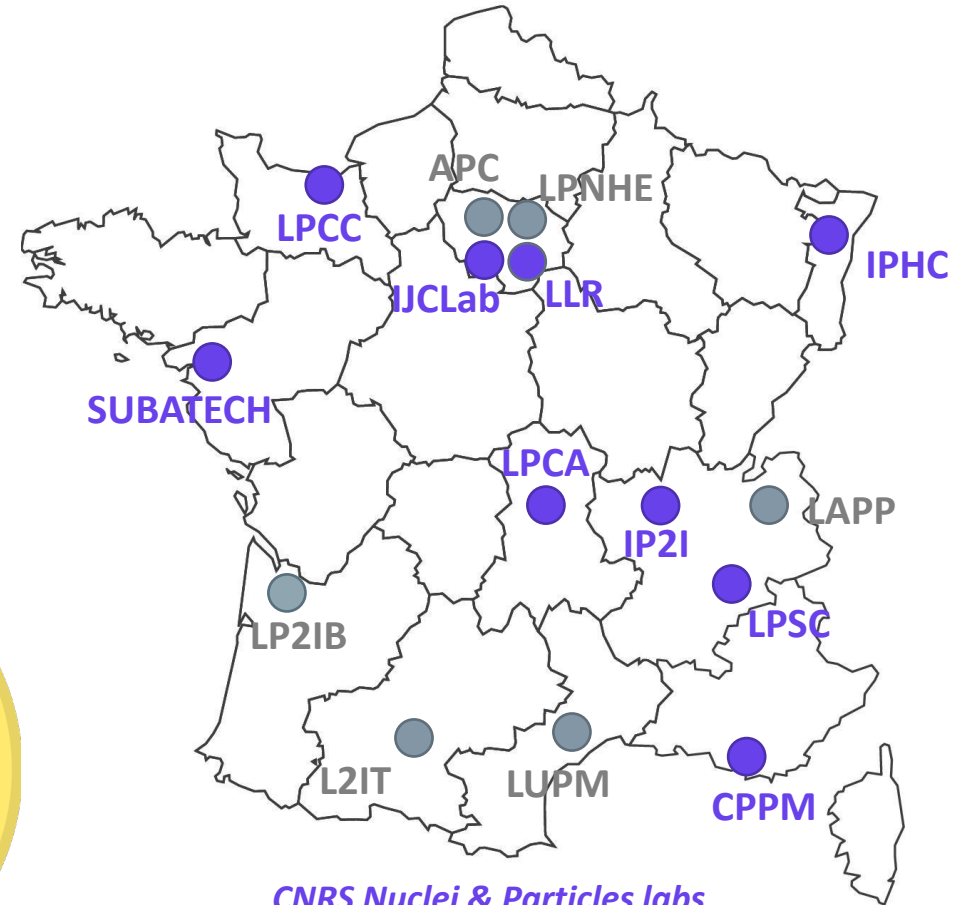
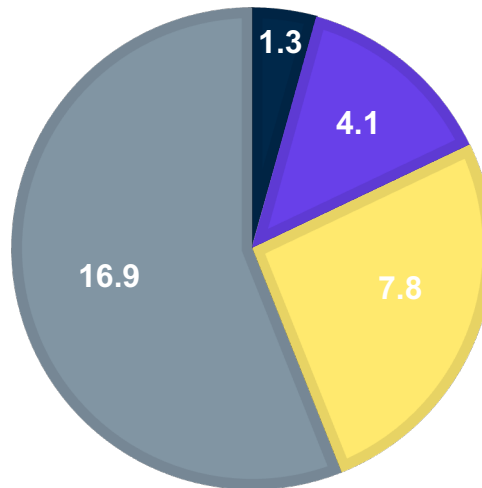
Stéphane Higuere

Hadrontherapy at IN2P3: key figures

9 IN2P3 labs involved
30 FTE/year @in2p3 + other collaborating institutes
90 persons involved (researchers and engineers)
19 subprojects

FTE/YEAR AT IN2P3

- Modelling
- Biological data
- Physico-chemical data
- Detector development

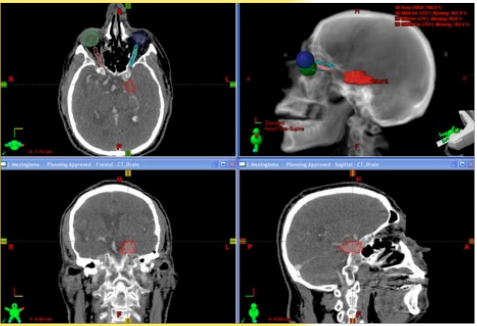


*CNRS Nuclei & Particles labs
involved in Hadrontherapy applications*

Research in Hadrontherapy

Imaging

Define tumoral/targeted
volume



Treatment planning system (TPS)



Treatment



Dose calculation model

Physical
processes

Biological
processes

Chemical
processes

Irradiation control

Beam

Range/dose
on-line

Dose QA

**Final goal: improve
hadrontherapy
treatments**

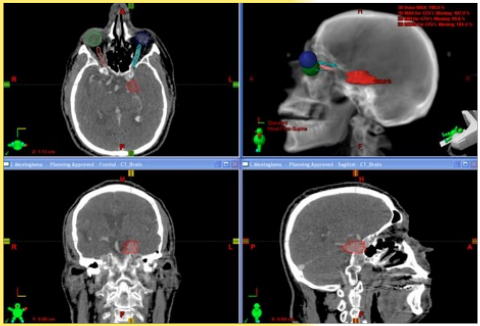
Fundamental research

Translational research

Research in Hadrontherapy

Imaging

Define tumoral/targeted
volume



Treatment planning system (TPS)



Treatment



Dose calculation model

Physical
processes

Chemical
processes

Biological
processes

Axis 1: Fundamental research

Irradiation control

Beam

Dose

Range

Axis 2: Translational research

- WP1
- WP2
- WP4
- WP3



NUCLÉAIRE
& PARTICULES

WP1
Multiscale,
multidisciplinary
modelling of radiation
effects

Coordinator:
Juliette Thariat

Task 1.1 Nano- and micro-metric maps of energy deposition and free radicals (Coord. E. Testa, IP2I)

Related projects: NanOx/GATE

Task 1.2 Biological and predictive models of tumour growth (Coord. M. Badoual, IJCLab)

Related projects: MODERATO

Task 1.3 Clinical data-driven modelling of NTCP and radio-resistance (Coord. J. Thariat, LPCC)

Related projects: aRTwin

3 sub-
projects

WP2
Measurement of nuclear
and physico-chemical
data

Coordinator:
Marie Vanstalle

Task 2.1 Nuclear data measurements (Coord. M. Vanstalle, IPHC)

Related projects: CLINM, FOOT

Task 2.2 Chemical data measurements (Coord. Q. Raffy, IPHC)

Related projects: Radiolysis

3 sub-
projects

WP3
Control of irradiation
delivery

Coordinator:
Sara Marcatili

Task 3.1 Beam monitoring (Coord. M-L. Gallin-Martel, LPSC)

Related projects: PEPITES@CNAO, N2 Beam profiler, Diamond Hodoscope, MATRIX

Task 3.2 Dose delivery characterisation (Coord. A-M. Frelin, GANIL)

Related projects: SCICOPRO, Micro-scintillator

Task 3.3 Range monitoring and tissue composition (Coord. S. Marcatili, LPSC)

Related projects: CLaRyS-S2C2, TIARA, XEMIS, vpgTLE

10 sub-
projects

WP4
Hadron interactions with
biological targets: from
cells to patients

Coordinator:
Claire Rodriguez-Lafrasse

Task 4.1 Production of biological data for simulation models (Coord. A-S. Wozny, IP2I)

Related projects: MODERATO, SIMUBIO

Task 4.2 Biological advantages of ions on radio-resistant and hypoxic tumours (Coord. C. Rodriguez-Lafrasse, IP2I)

Related projects: BIOHADRON

Task 4.3 Combinatorial approaches using ion therapy and other treatments (Coord. F. Chevalier, CIMAP)

Related projects: COMBION

3 sub-
projects

WP1. Multi-scale, multi-disciplinary modelling of radiation effects

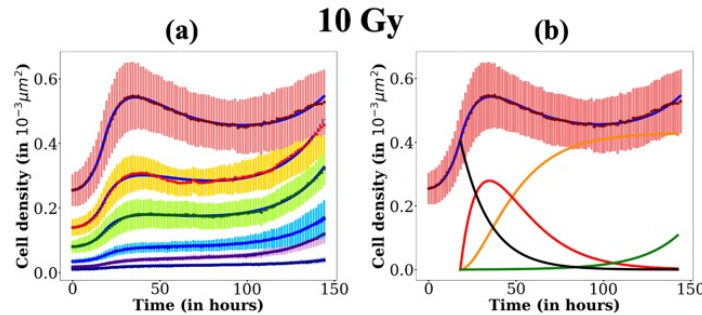
B. Dais's talk

MODERATO

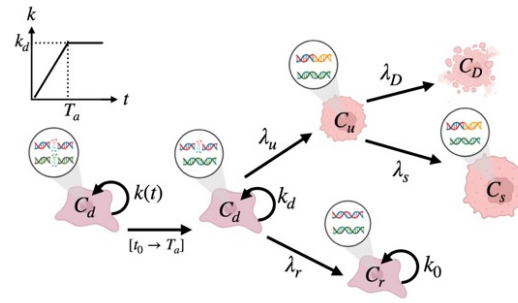
Beyond the Linear Quadratic model: introducing time and space dependency.



- Development of compartment model to access different cell populations (damaged, repaired, senescent cells, etc ...).
- Multi-parametric fit of cell culture data.



Experiment: points with error bars
Model: solid curves
 C_d : damaged cells
 C_u : unrepaired cells
 C_r : repaired cells
 C_s : senescent cells
 $C = C_d + C_u + C_r + C_s$ (Total cell population)



NanOx/ GATE

NanOx is a model for predicting cellular and biological doses based on nanometric and micrometric maps of energy deposits and free radicals distributions.

- Development of BioDoseActor for HT applications in GATE.
- Computational speed-up for future integration into TPS.

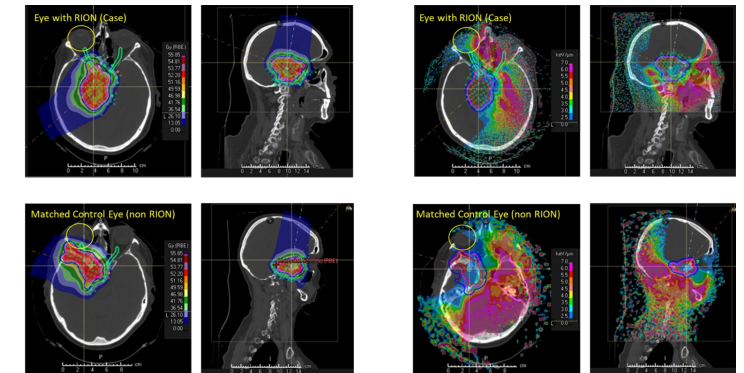


aRTwin



Experimental and clinical data-driven modelling of normal tissue toxicity and tumour radio-resistance.

- Extended clinical database: structural and functional assessment with toxicity classification.
- Real photons and protons plans + simulated ion plans.
- Development of NTCP data driven (ML) model.



Final goal: Integrated multi-ion, patient specific digital twin to simulate radiation effects under varying treatment conditions.

WP1. Multi-scale, multi-disciplinary modelling of radiation effects

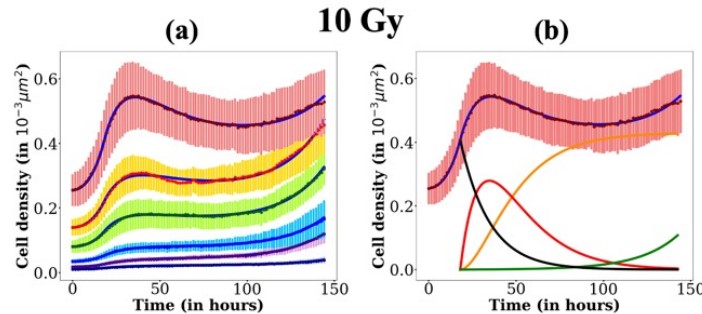
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MODERATO

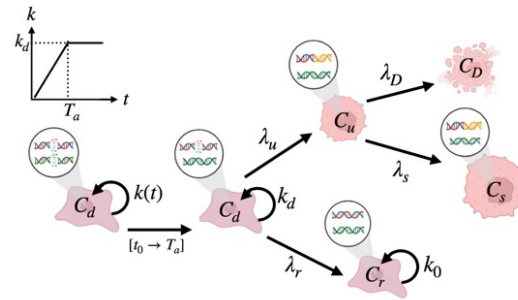
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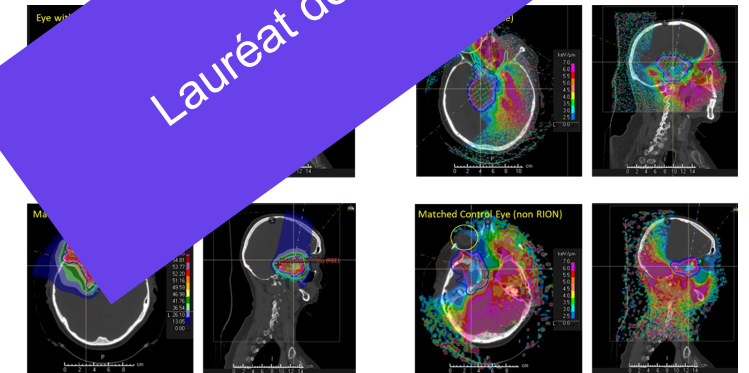
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aRTwin

Experimental and clinical data-driven modelling of normal tissue toxicity and patient radio-resistance.

- Extended clinical database and functional assessment with toxicological models.
- Real photons and particles dose and ion plans.
- Development of a multi-scale (molecular to organ) model.



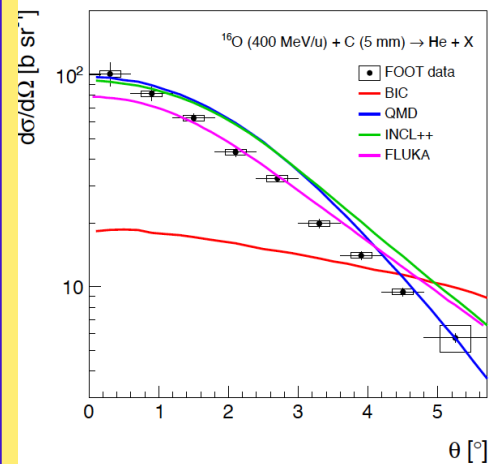
Final goal: Integrated multi-ion, patient specific digital twin to simulate radiation effects under varying treatment conditions.

WP2. Measurement of nuclear and physico-chemical data

- Measurement of nuclear cross section to improve dose calculation models
- Experimental validation of Geant4-DNA

FOOT

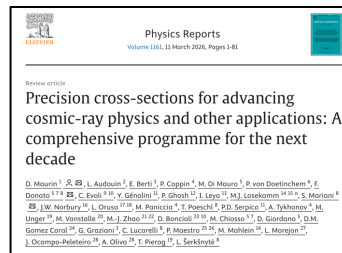
Measuring double differential cross-section for hadrontherapy with 5% accuracy



Multi-detector setup
International
collaboration with INFN
laboratories

- Première publication de sections efficaces différentielles $^{16}\text{O} \rightarrow ^{12}\text{C}$ (5mm) @ 400 MeV/u. R. Ridolfi et al., Phys. Rev. C **112**, (2025) **014610**

- Publication d'une feuille de route sur les mesures de sections efficaces de la cosmologie jusqu'aux applications (FOOT & CLINM notamment) dans Physics Reports : Maurin et al., 2026.

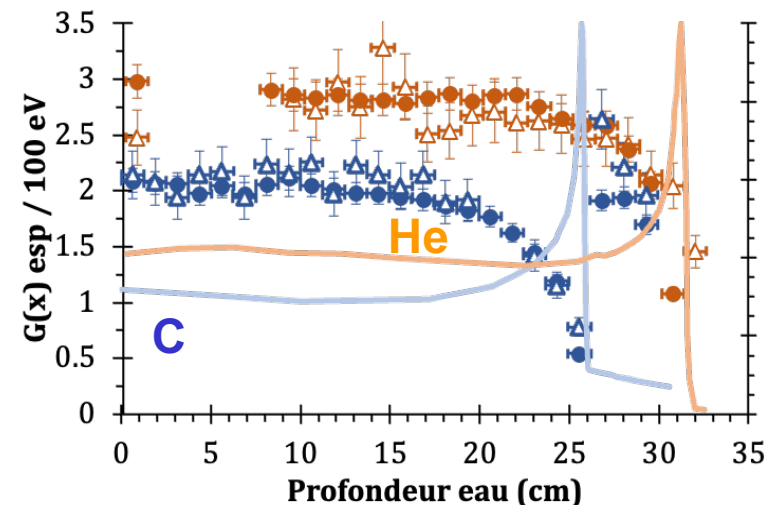


Radiolysis and CLINM

Q. Raffy's talk

- Study of LET effects, dose rate effects, oxygen effects on radiolysis of biomolecules + comparison to simulations.
- Study of radiolysis products in the fragments area (after the Bragg peak) and impact of secondary particles on indirect effects.

⇒ Measurement of yields of radiolysis species



WP4. Hadron interactions with biological targets: from cells to patient

Standardised data to support simulation models

Understand molecular and cellular mechanism responsible for higher RBE of ions

SIMUBIO & BioHADRON

C. Rodriguez
Lafrasse's talk

Characterization of the biological responses of tumoral cells (2D and 3D models) and pre-clinical models (in-ovo or in-vivo irradiations) to ions (protons, heliums, carbons).

- Production of biological data for simulation models
- Characterization of the biological advantages of ions in radioresistant and hypoxic tumors: study of nuclear and cytosolic DNA, mitochondrial parameters, proteostasis, cellular signalling, immune activation and cell death



COMBION

Combinatorial approaches using ion therapy alongside other treatments, studied in-cellulo, in-ovo & in-vivo.

- Inhibitors of proliferation pathways (focus on KRAS inhibitors)
- Proteasome inhibitors
- DNA repair inhibitors



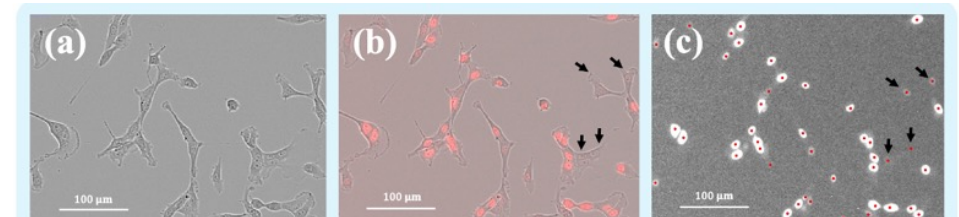
MODERATO

B. Dais's talk

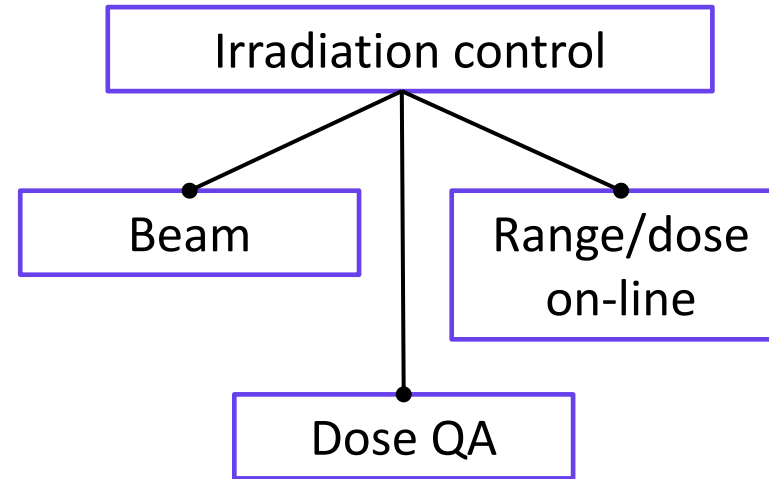
Multi-scale biophysical modeling of the effects of irradiation on cells in their microenvironment



- Cell culture time-lapse imaging (10 days)
- Different environments (i.e. cell density, hypoxia, nanoparticles...)
- Protons, ions and X-rays
- Development of a tracking software to analyse the fate of individual cells



WP3. Control of irradiation delivery



Essential elements of hadrontherapy optimisation:

- Feed TPS with accurate physical data (beam+dose)
- Allow quantification in preclinical experiments (beam+dose)
- On-line monitoring and eventually feedback to TPS (beam+range)

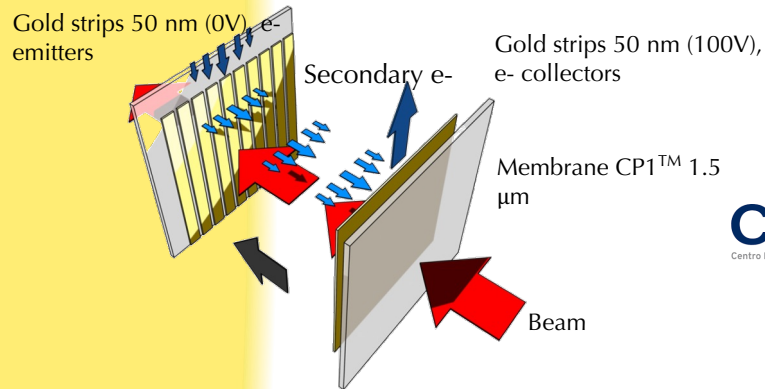
WP3. Control of irradiation delivery: beam monitoring

Non perturbative detectors

PEPITES @ CNAO

C. Thiebaux's talk

Nearly transparent (10->5 nm WET) distant (few m upstream) detector based on SEE.

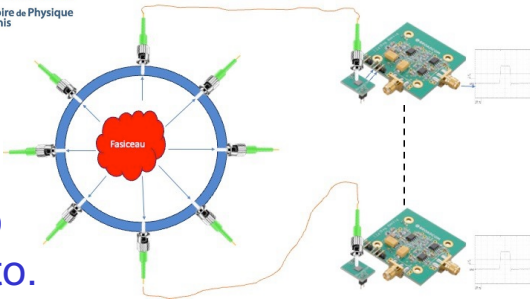


N2 Beam Profiler



L. Hardoin's talk

Transparent detector based on air as scintillating medium. Conceived for in-vitro experiments at BioAlto.



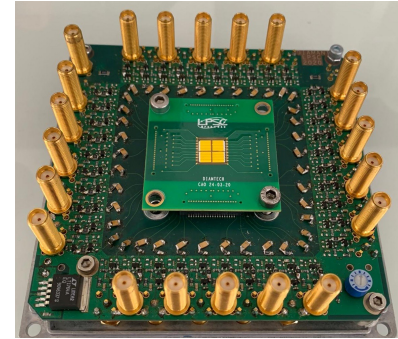
Fast detectors with large dynamic range (& FLASH applications)

Diamond Hodoscope



Many diamond-based detectors developed in the team. Hadrontherapy dedicated prototype focuses on:

- timing (~ 100 ps for single –nearly – MIPs)
 - compatibility with high count rates (100 MHz).
- + radiation resistant

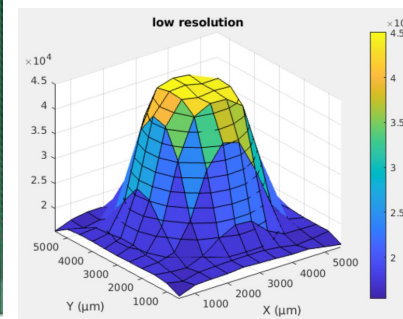
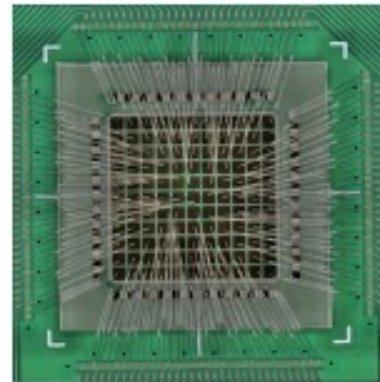


75 ps RMS time resolution

MATRIX

GaN based detector for real-time beam imaging.

- Temporal structure analysis for FLASH.
 - Sensitivity below 1 pA for radiolysis studies.
- + can be grown in large areas



- Current prototype: 11x 11 GaN PIN diodes on Sapphire substrate
- Goal: 128x128 matrix



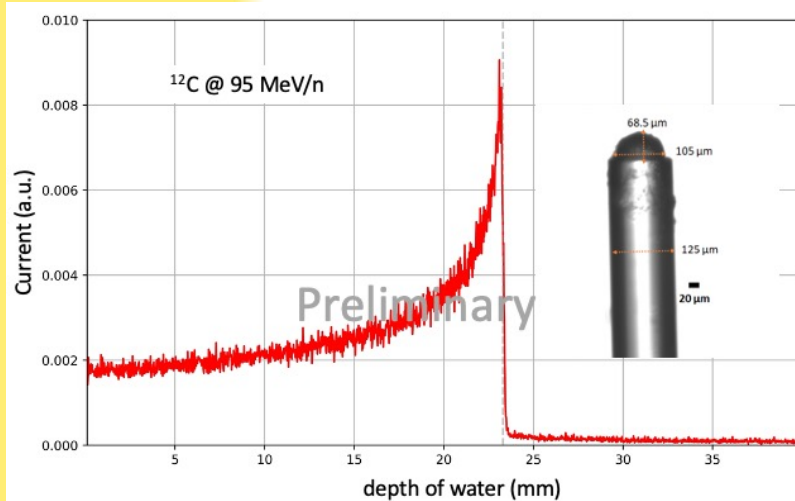
WP3. Control of irradiation delivery: dosimetry

Scintillation detectors as an alternative to Ionization Chambers

Micro-scintillator



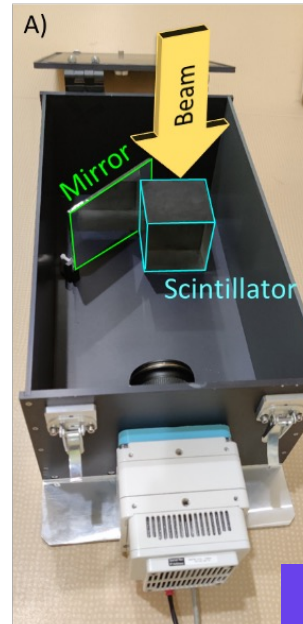
Micro-scintillator (inorganic) for LET and micro-dosimetry measurement.



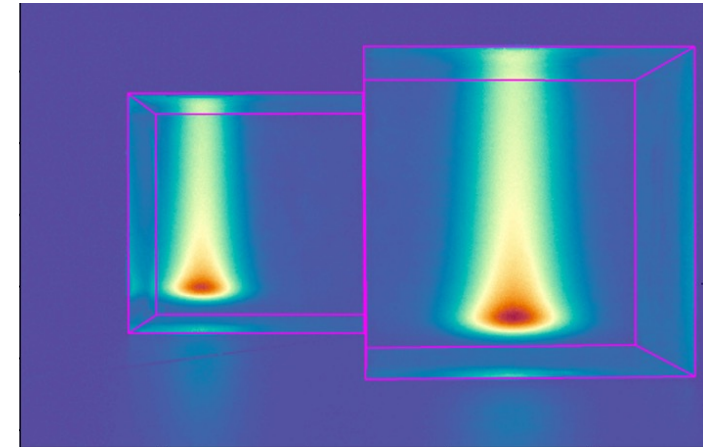
SCICOPRO



3D spatio-temporal dose deposition in PBS and its correlation with biological effects. Detector based on a monolithic organic scintillator cube.



- PBS image in a single acquisition
- Very high spatial resolution



2027: Study of AI methods in optical biases correction and reconstruction - TwinRISE European Project

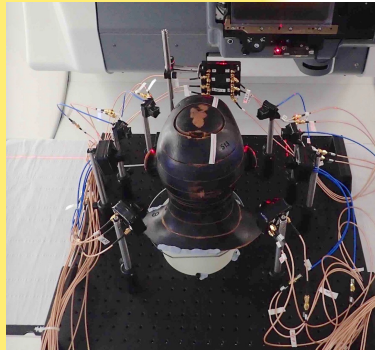
WP3. Control of irradiation delivery: on-line monitoring

TIARA

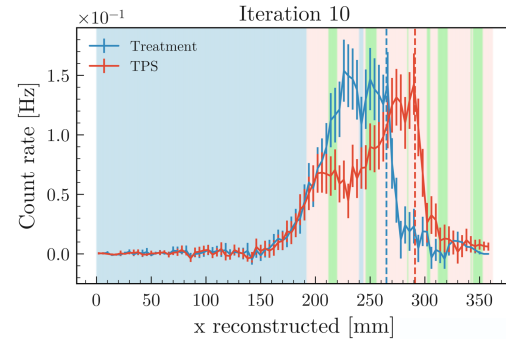
C. Abid's talk



From PG TOF distribution
to PG vertex distribution



Detection of anatomical
variations with the TIARA setup



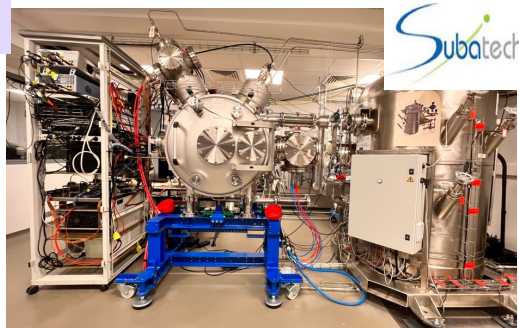
vpgTLE

Sinergy with TIARA for the fast
simulation of PGs in GATE



XEMIS2

Y. Ramsi's talk



3 γ imaging detector based
on liquid Xe technology

MC study for feasibility as a
range monitoring device

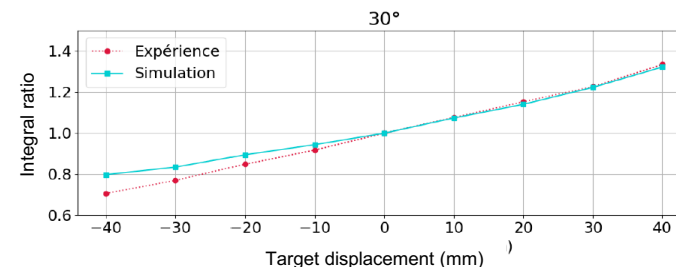
CLaRyS
S2C2

Energy integration of secondary
radiation with high intensity pulsed
beams from CAL-Proteus One

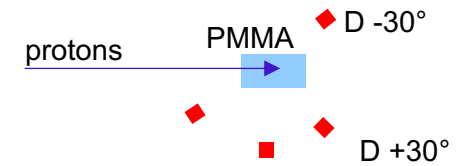


PbWO4 scintillators+ fast PMTs. ASIC QDCs developed at LPSC

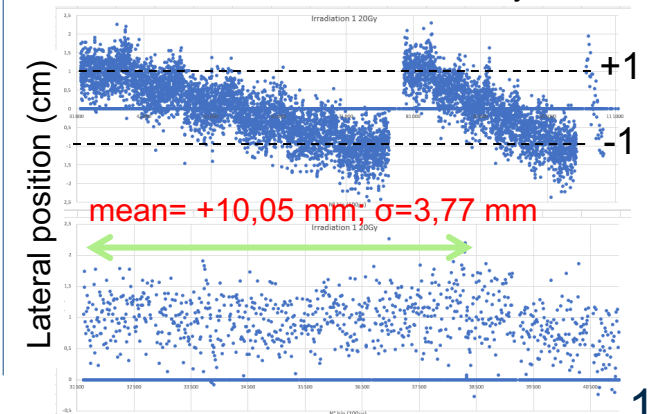
68 MeV protons at ARRONAX
(longitudinal target displacement)



140 MeV protons at P1 - CAL
(transverse target displacement)

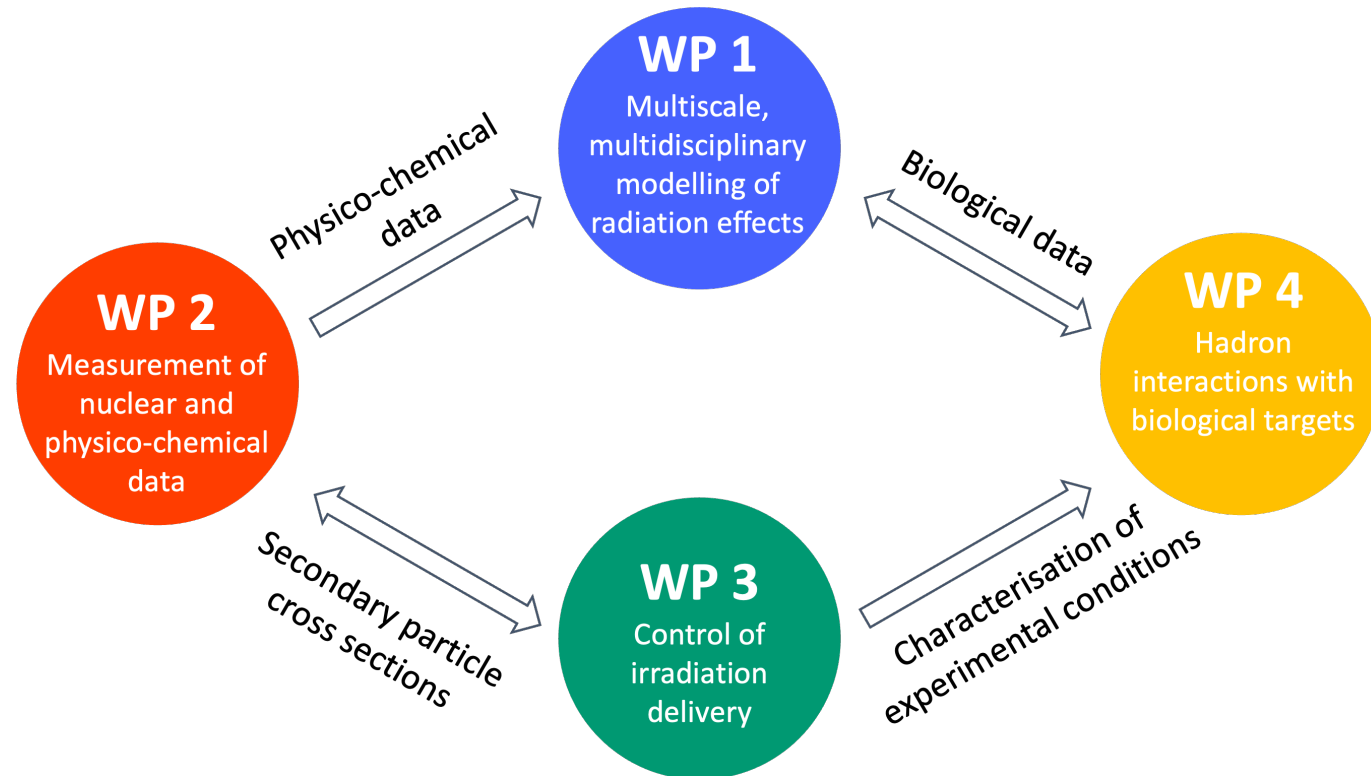


Irradiation 2x2 cm² 20 Gy



Conclusions and perspectives

- **MP récent, construit sur des projets existants à l'IN2P3**
 - ❖ Financements souvent mono-équipe sur AaP
 - ❖ Collaboration hors IN2P3 existantes
- ❖ Lieu d'animation pour encourager les collaborations inter-équipes (réponse à AaP communes)
- ❖ Proposition: Workshop en ligne en Septembre (evento dans les prochains jours)



Conclusions and perspectives

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- **Projets très dépendants du temps faisceau**
 - ❖ Accord IN2P3 – **CAL** arrivé à terme
 - ❖ Accord IN2P3 – **CNAO** : renouvellement et renforcement des collaborations existantes
 - Beam/range monitoring – PG imaging (**PEPITES**, TIARA)
 - Radiobiology (**BioHadron**)
 - TPS – radiolysis (**CLINM**)
 - BNCT – neutrons measurements (to be discussed)
 - ❖ Accord IN2P3 - **CYCLHAD**
- **Accord cadre IN2P3 – INFN** signé en 2026: opportunité pour des futures collaborations
- Premier **poste de CR** depuis 2016 sur les activités inter-disciplinaires

MP Hadrontherapy at the Particle Therapy Co-Operative Group (PTCOG) conference



Michael Goitein Awards, Biology
Claire Rodriguez-Lafrasse





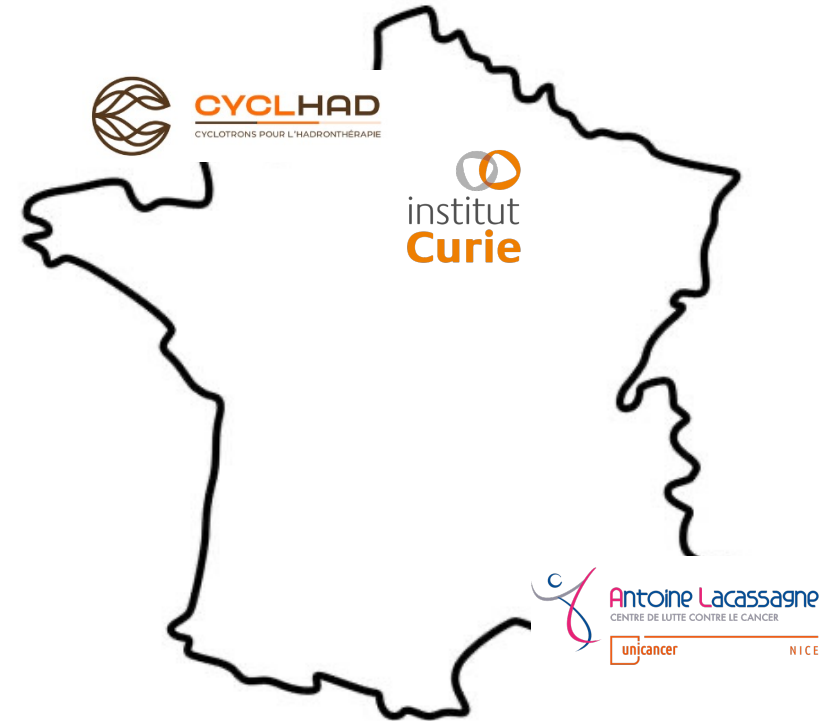
NUCLÉAIRE
& PARTICULES

Additional slides

Irradiation facilities



Main platforms & collaborations



French Hadrontherapy facilities

Current collaborations

Multi-scale modelling

French:
CREATIS, LIRIS, TIMC, ISMO, CFB

International: GATE collaboration

Platforms: CPO, RadEx, BioAlto

Physico-chemical data

French:
Icube, IBMP, Icans, Aerial-CRT

International: CNAO, CNRS, QST-Himac, **INFN**, Geant4 collaboration

Platforms: CAL, CNAO, GANIL, CYRCE, GSI, HIMAC...

**HT
project**

Irradiation control

French:
IP Paris, DIAMFAB, CHREA, CLCC, CINAM, CREATIS, ARRONAX

International: CNAO, Ruhr University, WGPTC, Roma Tor Vergata

Platforms: CAL, CNAO, GANIL, ARRONAX, CYCLHAD, BioAlto...

Biological data

French:
ARRONAX, IAB, CEA, CLCC

International: CNAO, NIRS, IFIN-HH

Platforms:
CAL, CNAO, ARRONAX, GANIL

Project name	Primary IN2P3 lab	PI
Nanox	IP2I Lyon	E.Testa/M. Beuve
MODERATO	IJCLAB Orsay (Paris)	M. Badoual
aRTwin	LPCC Caen	J. Thariat
CLINM	IPHC Strasbourg	M. Vanstalle
FOOT	IPHC Strasbourg	C. Finck
Radiolysis	IPHC Strasbourg	Q. Raffy
PEPITES	LLR Palaiseau (Paris)	M. Verderi
N2 Beam Profiler	IJCLAB Orsay (Paris)	Q. Mouchard
Diamond hodoscope	LPSC Grenoble	M-L. Gallin-Martel
MATRIX	IPHC Strasbourg	S. Higuieret
SCICOPRO	GANIL Caen	A-M. Frelin
Micro-scintillator	LPCC Caen	M. Rousseau
CLaRyS-S2C2	LPSC Grenoble	D. Dauvergne
TIARA	LPSC Grenoble	S. Marcatili
XEMIS/SMART	SUBATECH Nantes	N. Beaupere
vpgTLE	IP2I Lyon	E. Testa
SIMUBIO	IP2I Lyon	A-S. Wozny
BioHADRON	IP2I Lyon	C. Rodirguez-Lafrasse
COMBION	CIMAP Caen	F. Chevalier

WP synergies

