

Development of the analytical model RAMBI of irradiation beamline

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International Conference on 3D dosimetry (IC3DDose), June 2026, Caen



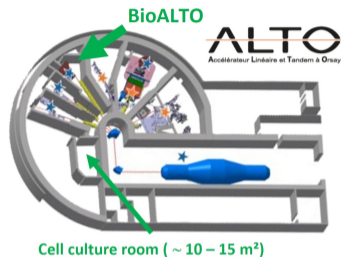
“BioALTO” Projet: “Hadron biology” Platform @ ALTO

Objective

- Development of a platform dedicated to **radiobiological research for therapies involving ions** (hadrontherapy, Targeted Alpha Therapy, Boron Neutron Capture Therapy)
- **BioALTO beamline based on Radiograaff device** (previously at the IP2I 4 MV accelerator in Lyon)

Characteristics of ALTO

- 14.5 MV Tandem “Van de Graaff” accelerator
- **Wide range of intense (up to μA) ions beams ($^1\text{H} \rightarrow ^{79}\text{Au}$)**
- Continuous beam
- Additional pulsed beam possible (100 ns to 100 μs)



Part.	Energies [MeV] (MeV/u)	Ion range in water [μm]*
^1H	4 – 25 (4 – 25)	160 – 6190
^4He	15 – 43 (3.8 – 10.8)	130 – 1290
^7Li	50 (7.1)	480
^{12}C	87.5 (7.29)	170
^{16}O	128 (8)	140

*(at 5 cm air)



Project funded by the Ile de France Region

“BioALTO” Project: Main tasks

Instrumentation

- Upgrade the Radiograaff device to exploit the wide range of ions
- Development of new beam diagnostics and dosimetry tools
 - ▶ In line: Diamond & scintillating fibre counters
 - ▶ Exit: Microdosimeters and beam monitoring profiler based on air fluorescence

Biology

- Equipment of a cell culture room nearby

⇒ Access to external users → platform in 2-3 years

Modeling

- Digital twin (Monte Carlo simulations): reference calculations
- Analytical model: optimization and operation of the beamline



Fig. 1: BioALTO beamline under installation

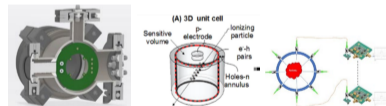


Fig. 2: From left to right: Diamond counter, Si microdosimeter, beam profiler based on air fluorescence

The 2 “collimated-diffusor” design

Specifications

- Diameter of the irradiation field (sample): 15 mm
- Dose homogeneity: $\epsilon_{\phi_4} = 2\%$
- Dose rate: \sim Gy/min (\Rightarrow Beam intensity of $\sim 0.1 - 1$ pA)

Accelerators for nuclear physics

- Examples: 4 MV @ IP2I, BioALTO
- Uncertainties on beam direction ($1^\circ \rightarrow 17$ mm at 1 m)
- Nominal beam intensities > 1 nA

Solution: 2 “collimated diffusors” (CD)

- Single CD: sensitivity to incident beam direction
- 2 CD:
 - ▶ Outcoming particles aligned with the beam line
 - ▶ Typical factor of beam intensity reduction: 100 – 1000 for $\theta_0 = 0$

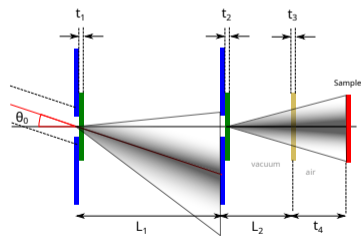
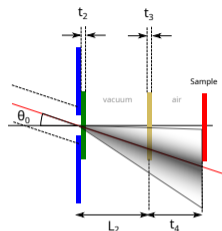
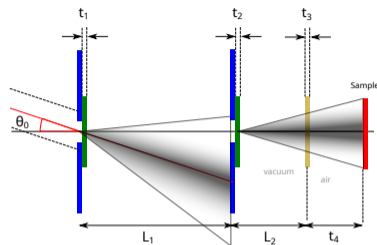


Fig. 3: Top: Single CD design sensitive to incident beam direction uncertainties: bottom: 2 CD design.

Objectives of the analytical model

2 modes

- “Inverse mode”: Optimization of the **diffusors thicknesses** and determination of the required **incident beam intensity**
- “Direct mode”: Prediction of the flux distribution and intensity at beamline exit



General input data	Specific data	Inverse	Direct
Beam radius R_0	Input	Energy E_4	Energy E_0
Angle θ_0		Intensity I_4 (dose rate)	Intensity I_0
L_1, L_2, t_3, t_4	Output	Homogeneity $\epsilon_\phi = 2\%$	Thicknesses t_1, t_2
Collimator radius R_1, R_2		Thicknesses t_1, t_2	Energy E_4
Collimator material (gold)		Energy E_0	Flux $\Phi_4(r_4)$ ($I_4, \sigma_{r,4}$)
Sample radius R_4		Intensity I_0	

Beam diffusion model: Gaussian model based on MC simulation

Parametrization of A_Ω and σ_Ω

$$\frac{dP}{d\Omega}(E, Z, t) = A_\Omega(E, Z, t) \exp\left(-\frac{1}{2} \frac{r^2}{\sigma_\Omega(E, Z, t)^2}\right)$$

- where, at first order:

$$\sigma_\Omega(E, Z, t) \propto \frac{\sqrt{t}}{E} \quad \text{and} \quad A_\Omega(E, Z, t) \propto \frac{E}{t}$$

Relationship between angular and spatial distrib.

$$\sigma_r = \sigma_\Omega L \quad \text{and} \quad A_r = \frac{A_\Omega}{L^2}$$

Simulated geometry (GATE v9.3)

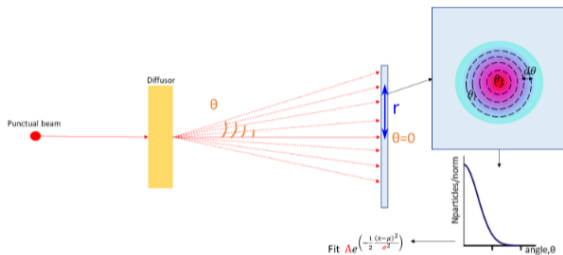


Fig. 4: Simulated geometry for obtaining the parameters of the Gaussian distribution

Advantage of MC sim. wrt Moliere's theory¹

- Consistency of the results with the MC simulation used to validate the model

¹ (H. A. Bethe [Mar. 1953]. "Molière's Theory of Multiple Scattering". In: *Physical Review* 89 [6], p. 1256. DOI: 10.1103/PhysRev.89.1256)

Beam diffusion model: Results and limit of validity

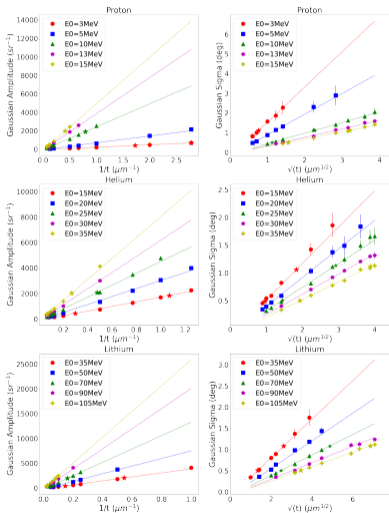


Fig. 5: A_{Ω} and σ_{Ω} values obtained with MC simulations.

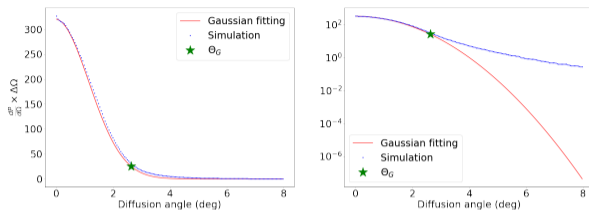


Fig. 6: Illustration of the limit of validity of the Gaussian model (10 MeV) α particles on 2 μm gold foil.

Conclusions

- A_{Ω} and σ_{Ω} vary according the expected scaling laws (slight deviations taken into account in additional terms in the parametrization)
- Limit of validity: θ_G where the deviation between simulated and Gaussian distrib. equal to 10%

Criteria of validity of the beamline model: Validity of the beam transport

Critical angles θ_C

$$\theta_{C,C2} = \theta_0 + \frac{R_2}{L_1} < \theta_G$$

et
$$\theta_{C,C4} = \frac{R_4}{L_2 + t_3 + t_4} < \theta_G$$

Constraint on the minimal foil thicknesses (t_{min})

One can show that:

$$\theta_G = C_G \frac{Z}{E} \sqrt{t}$$
$$\Rightarrow t_{min} = \frac{\theta_C^2 E^2}{C_G Z^2}$$

where C_G is a proportionality coefficient

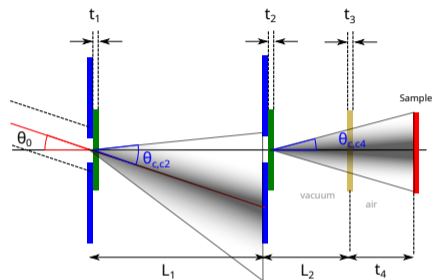


Fig. 7: Definition of the critical angles $\theta_{C,C2}$ and $\theta_{C,C4}$ to properly model the beam transport.

Criteria of validity of the beamline model: Other criteria

Hole-size criterion

- Extension of collimator's hole (radius R) taken into account by means of **convolution** of the calculated distributions with Gaussians of $\sigma = \sqrt{2R}$

$$\sigma_{\Omega,1,conv}^2 = \sigma_{\Omega,1}^2 + \left(\frac{R_1}{\sqrt{2}L_1} \right)^2$$

$$\sigma_{\Omega,2,conv}^2 = \sigma_{\Omega,2}^2 + \left(\frac{R_2}{\sqrt{2}(L_2 + t_3 + t_4)} \right)^2$$

$$\Rightarrow \frac{R_1}{\sigma_{r,2}} \ll 1 \text{ and } \frac{R_2}{\sigma_{r,4}} \ll 1$$

Energy-loss criterion

- Parametrization of **diffusion model** depends on ions' energy

\Rightarrow Ion's energy loss in the diffusors must be negligible \Rightarrow **Limit on the diffusor thickness t_{max}**

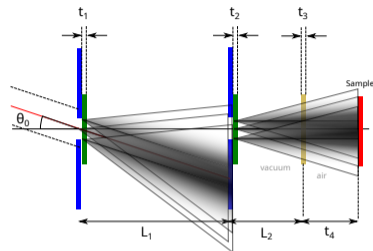


Fig. 8: Illustration of the convolution of the calculated distributions to take into account the extension of collimator's hole.

Modeling and optimization of the 2nd diffusor

(Gaussian) flux distribution at sample position

$$\Phi_4(r_4) \propto A_{r,4} \exp\left(-\frac{1}{2} \left(\frac{r_4}{\sigma_{r,4,conv}}\right)^2\right)$$

Homogeneity criterion (ϵ_{Φ_4})

One can show that:

$$\epsilon_{\Phi_4} = \frac{\sigma_{\phi_4}}{\phi_4(R_4)} \sim \frac{1}{\sqrt{48}} \left(\frac{R_4}{\sigma_{r,4,conv}}\right)^2$$

- Optimization for a given ion type and energies E_4 (sample) and E_3 (beam exit):

$$\epsilon_{\Phi_4} = 2\% \rightarrow \sigma_{r,4,conv} \xrightarrow{\text{Gaussian model}} t_2 \rightarrow E_1 \quad (\text{before 2}^{\text{nd}} \text{ diffusor})$$

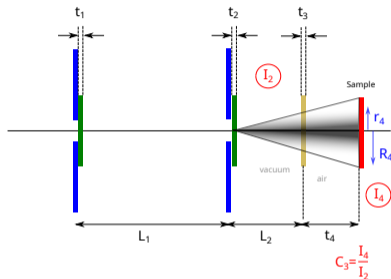


Fig. 9: Illustration of the transmission coefficient C_3 .

Transmission coeff. (C_3)

$$C_3 = a \left(\frac{R_4}{\sigma_{r,4,conv}}\right) A_{r,4} \pi R_4^2$$

Modeling and optimization of the 1st diffusor

Transmission coefficients of the collimators

- $C_1 = \frac{S_1(\theta_0)}{S_0} \sim \frac{S_1}{S_0}$
- $C_2 \sim \frac{dP_1}{d\Omega}(\theta_0)\Delta\Omega = A_{\Omega,1,conv} \exp\left(-\frac{1}{2}\left(\frac{\theta_0}{\sigma_{\Omega,1,conv}}\right)^2\right) \frac{S_2}{L_1^2}$

Optimization

- Use of the scaling laws of σ_{Ω} and A_{Ω} neglecting the impact of hole's extension

$$\frac{l_2}{l_0} = C_1 C_2 \propto \frac{1}{t_1} \exp\left(-\frac{1}{2}\left(\frac{\theta_0^2}{t_1 \times (g(Z,E))^2}\right)\right)$$

$$\Rightarrow l_2 \text{ max for } t_1 = \frac{\theta_0^2}{2(g(Z,E))^2}$$

where $g(Z,E)$ is obtained through the Gaussian model of diffusion

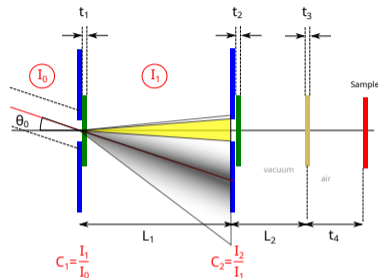


Fig. 10: Illustration of the transmission coeff. C_1 and C_2 .

Validation of the model against MC simulations: Transmission coeff. C_2 and C_3

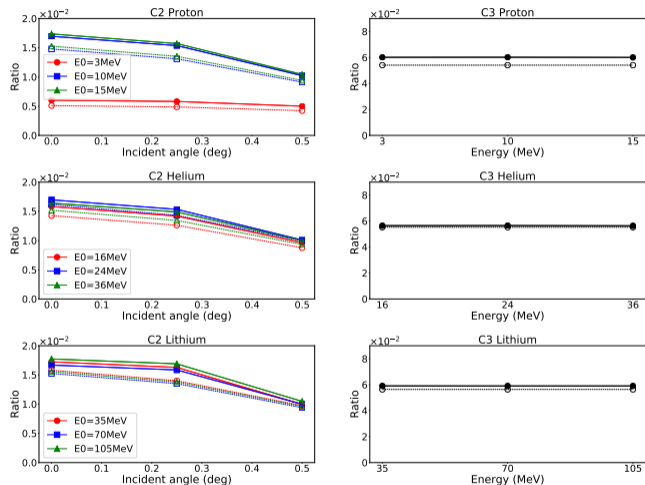


Fig. 11: Transmission coefficients obtained by MC simulations (full symbols) and analytical model (open symbols).

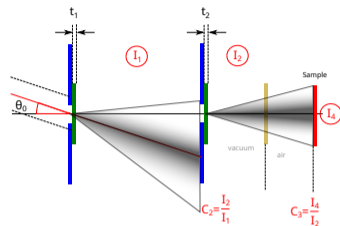


Fig. 12: Transmission coefficients C_2 and C_3 .

- Calculations with optimized ticknesses
- Agreement within $\sim 10\%$ between Monte Carlo simulations and the analytical model

Application of the model: Thicknesses of the diffusors

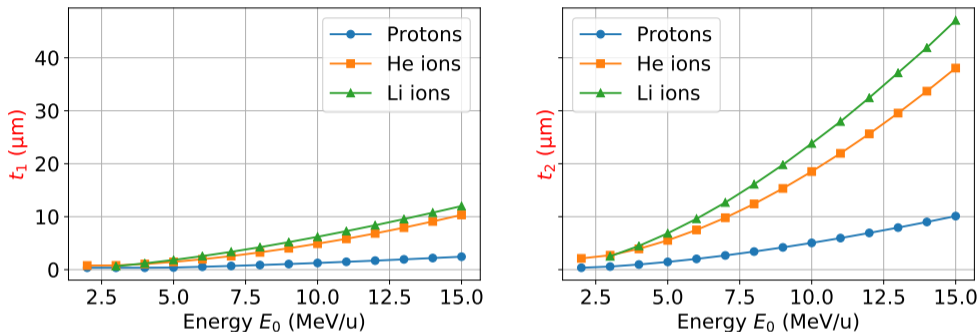


Fig. 13: Diffusor thicknesses t_1 and t_2 as a function of the incident beam energy E_0 with $\theta_0 = 0.5^\circ$.

Orders of magnitude

- 1st diffusor: a few μm
- 2nd diffusor: a few tens of μm

Trend as a function of beam energy

- Quadratic behavior consistent with $\sigma_\Omega(E, Z, t) \propto \frac{\sqrt{t}}{E}$

Application of the model: Beam current at the beamline entrance/exit

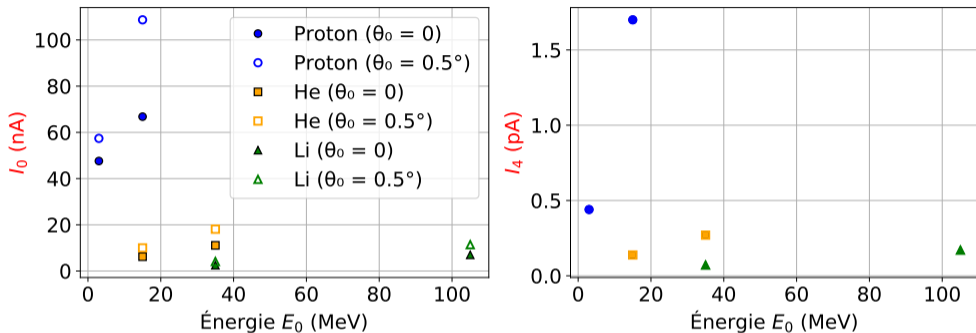


Fig. 14: Initial and final beam intensities required to achieve a dose rate of 2 Gy/min.

Orders of magnitude for a dose rate of 2 Gy/min

- Exit: a few tenth of pA
- Entrance: from a few nA (He, Li) to tens of nA (protons)

Impact of the incident beam angle θ_0

- $\sim 50\%$ between 0° and 0.5°

Conclusion

Development of the analytical model “RAMBI”

- Modeling of the 2 “collimated-diffusor” BioALTO beamline distribution and intensity at beamline exit
- “Inverse mode”: Optimization of the diffusors thicknesses and determination of the required incident beam intensity

Perspectives

- Application to the BioALTO platform
 - ▶ Wide range of ions and energies
 - ▶ Radioprotection issues (beam limitations \Rightarrow re-optimization)
- Potential application to new irradiation beamlines
 - ▶ Advantage of the 2 “collimated-diffusor” design: Robust wrt uncertainties in incident beam direction

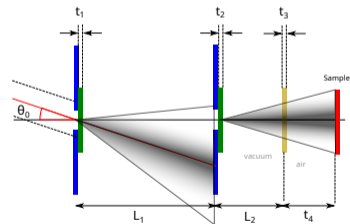


Fig. 15: BioALTO beamline under installation (top) and scheme (bottom).

Acknowledgments

Current collaborators of the “BioALTO” project

- **IJCLab**: Philippe Lanièce, Amélia Maia Leite, Quentin Mouchard, Louis Hardouin
- **IP2I**: M. Alcocer-Avila, M. Beuve, G. Garde, E. Testa, Y. Zoccarato
- **LPSC**: R. Delorme

Fundings



Thank you for your attention

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Constraint on the minimal foil thicknesses (t_{min})

$$\theta_G \propto \sqrt{N} = \sqrt{\frac{t}{\lambda}} \propto \sqrt{\sigma t} = C_G \frac{Z}{E} \sqrt{t}$$
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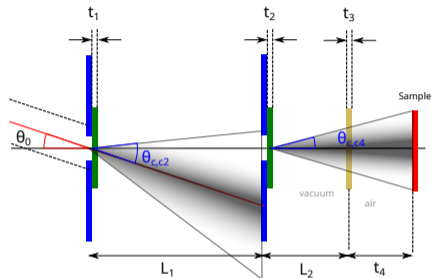


Fig. 16: Definition of the critical angles $\theta_{C,C2}$ and $\theta_{C,C4}$ to properly model the beam transport.

Modeling and optimization of the 2nd diffusor

Flux distribution at sample position

$$\Phi_4(r_4) = \left(\frac{I_0 C_1 C_2}{Ze} \right) A_{r,4} \exp \left(-\frac{1}{2} \left(\frac{r_4}{\sigma_{r,4,conv}} \right)^2 \right)$$

Homogeneity criterion (ϵ_{Φ_4})

One can show that:

$$\epsilon_{\Phi_4} = \frac{\sigma_{\Phi_4}}{\Phi_4(R_4)} = \left(\frac{a \left(\frac{\sqrt{2}R_4}{\sigma_{r,4,conv}} \right)}{\left[a \left(\frac{R_4}{\sigma_{r,4,conv}} \right) \right]^2} - 1 \right)^{\frac{1}{2}} \sim \frac{1}{\sqrt{48}} \left(\frac{R_4}{\sigma_{r,4,conv}} \right)^2$$

where $a(X) = \int_0^X x \exp \left(-\frac{1}{2}x^2 \right) dx = \frac{2}{X^2} \left(1 - \exp \left(-\frac{1}{2}X^2 \right) \right)$

- Optimization for a given ion type and energies E_4 (sample) and E_3 (beam exit):

$$\epsilon_{\Phi_4} = 2\% \rightarrow \sigma_{r,4,conv} \xrightarrow{\text{Gaussian model}} t_2 \rightarrow E_1 \quad (\text{before 2}^{\text{nd}} \text{ diffusor})$$

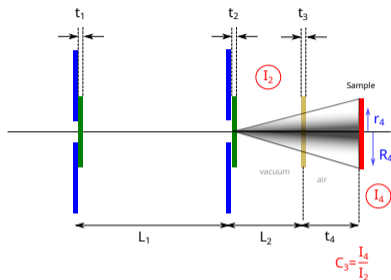


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