



UNIVERSITÀ  
DEGLI STUDI  
DI MILANO



# The Copper Trilogy: exploring $^{61}\text{Cu}$ , $^{64}\text{Cu}$ and $^{67}\text{Cu}$ at the Bern Medical Cyclotron

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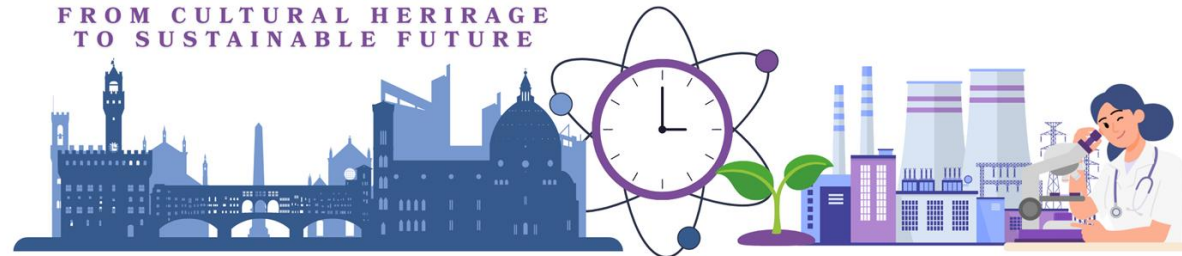
<sup>1</sup> Tera-Care Foundation, Cologny, Switzerland

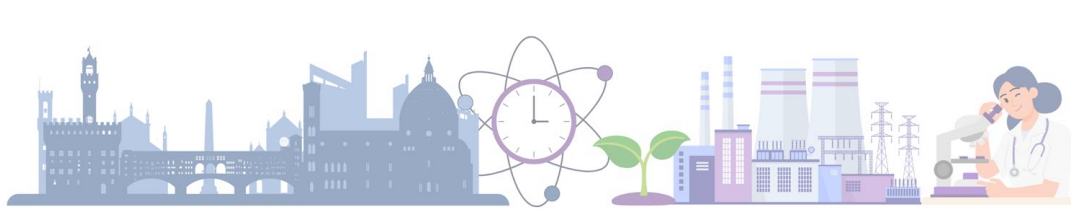
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**ISOTOPIC TIME MACHINE**  
FROM CULTURAL HERITAGE  
TO SUSTAINABLE FUTURE





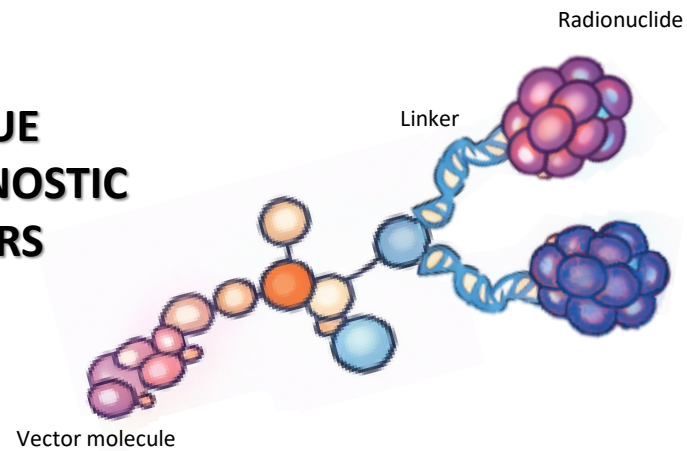
# Copper radioisotopes in nuclear medicine

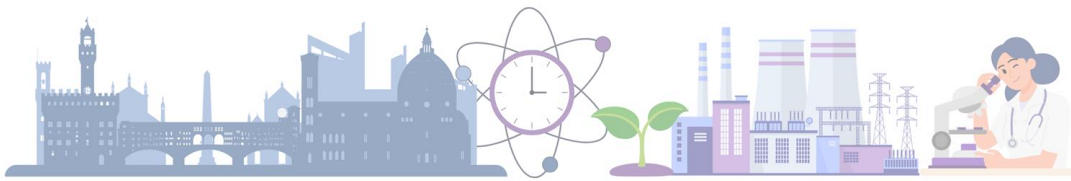
**Cu-61**  
 3.339 h  
 ec,β<sup>+</sup>: 100%

**Cu-64**  
 12.7006 h  
 ec,β<sup>+</sup>: 61.5%  
 β<sup>-</sup>: 38.5%

**Cu-67**  
 61.83 h  
 β<sup>-</sup>: 100%

**TRUE  
 THERANOSTIC  
 PAIRS**





# Production routes

## Cu-61

3.339 h  
 ec,β<sup>+</sup>: 100%

- **Low-energy protons\*:**  $^{61}\text{Ni}(p,n)^{61}\text{Cu}$
- Low-energy protons\*:  $^{64}\text{Zn}(p,\alpha)^{61}\text{Cu}$

## Cu-64

12.7006 h  
 ec,β<sup>+</sup>: 61.5%  
 β<sup>-</sup>: 38.5%

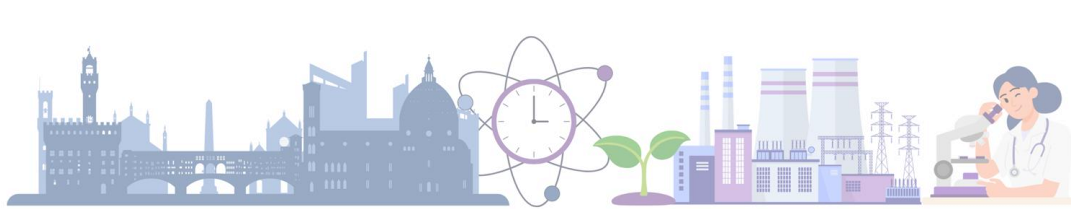
- **Low-energy protons\*:**  $^{64}\text{Ni}(p,n)^{64}\text{Cu}$
- **Neutrons:**  $^{64}\text{Zn}(n,p)^{64}\text{Cu}$ ; **c.a.**  $^{65}\text{Cu}(n,2n)^{64}\text{Cu}$
- Low-energy protons\*:  $^{67}\text{Zn}(p,\alpha)^{64}\text{Cu}$ ; **c.a.**  $^{65}\text{Cu}(p,pn)^{64}\text{Cu}$

## Cu-67

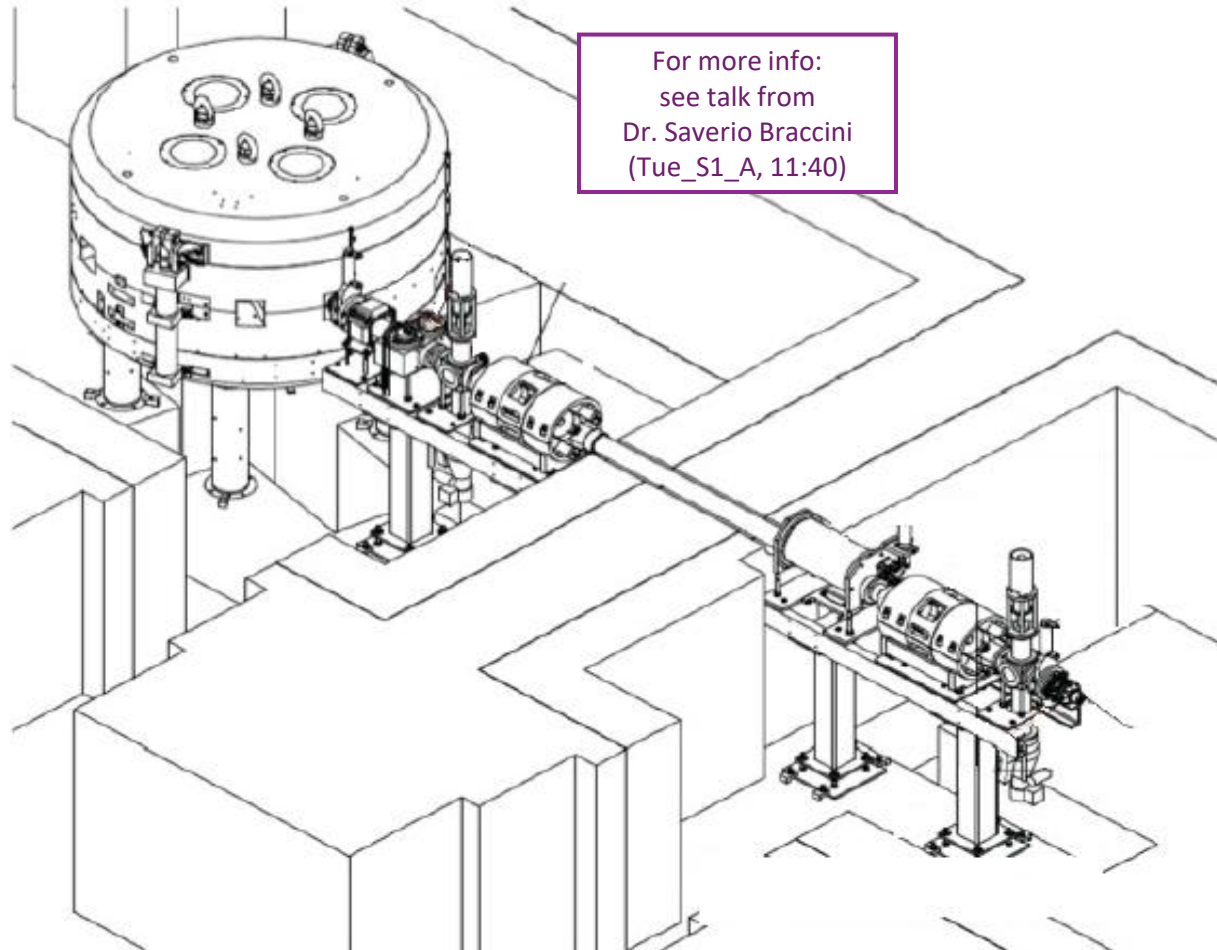
61.83 h  
 β<sup>-</sup>: 100%

- **Neutrons:**  $^{67}\text{Zn}(n,p)^{67}\text{Cu}$
- **Spallation process:**  $^{75}\text{As}(p,\text{spall})^{67}\text{Cu}$
- **High-energy protons:**  $^{68}\text{Zn}(p,2p)^{67}\text{Cu}$
- Low-energy protons\*:  $^{70}\text{Zn}(p,\alpha)^{67}\text{Cu}$

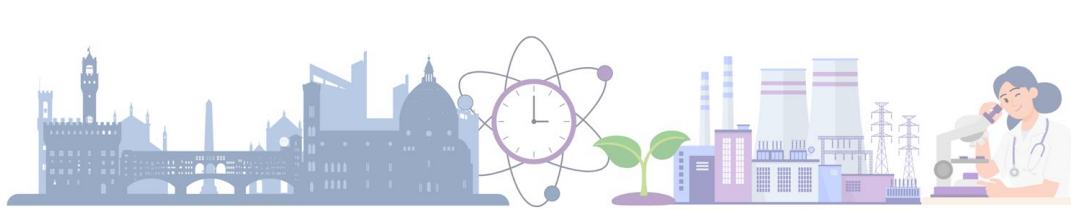
\*E<sub>p</sub> ≤ 20 MeV



# The Bern Medical Cyclotron laboratory

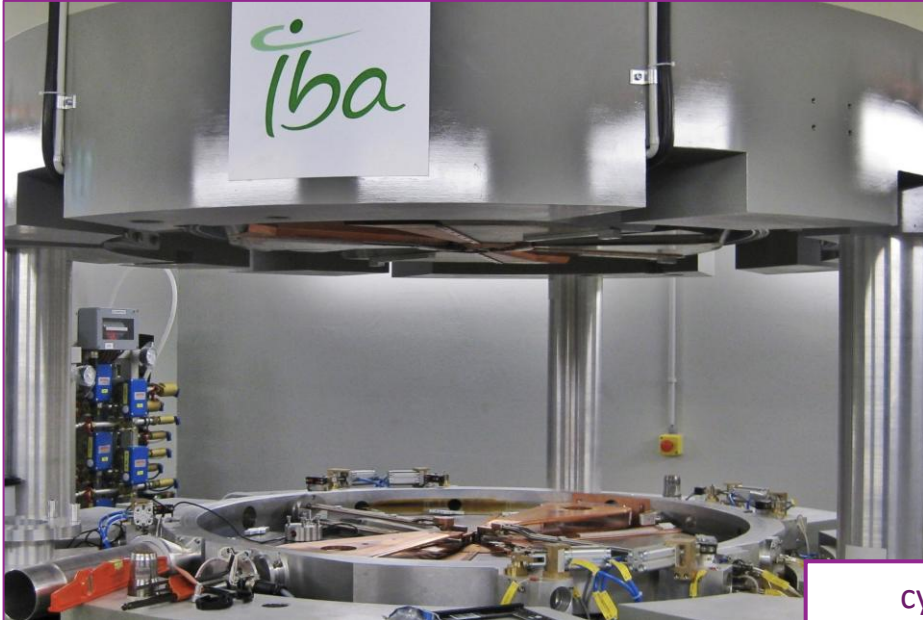


For more info:  
see talk from  
Dr. Saverio Braccini  
(Tue\_S1\_A, 11:40)



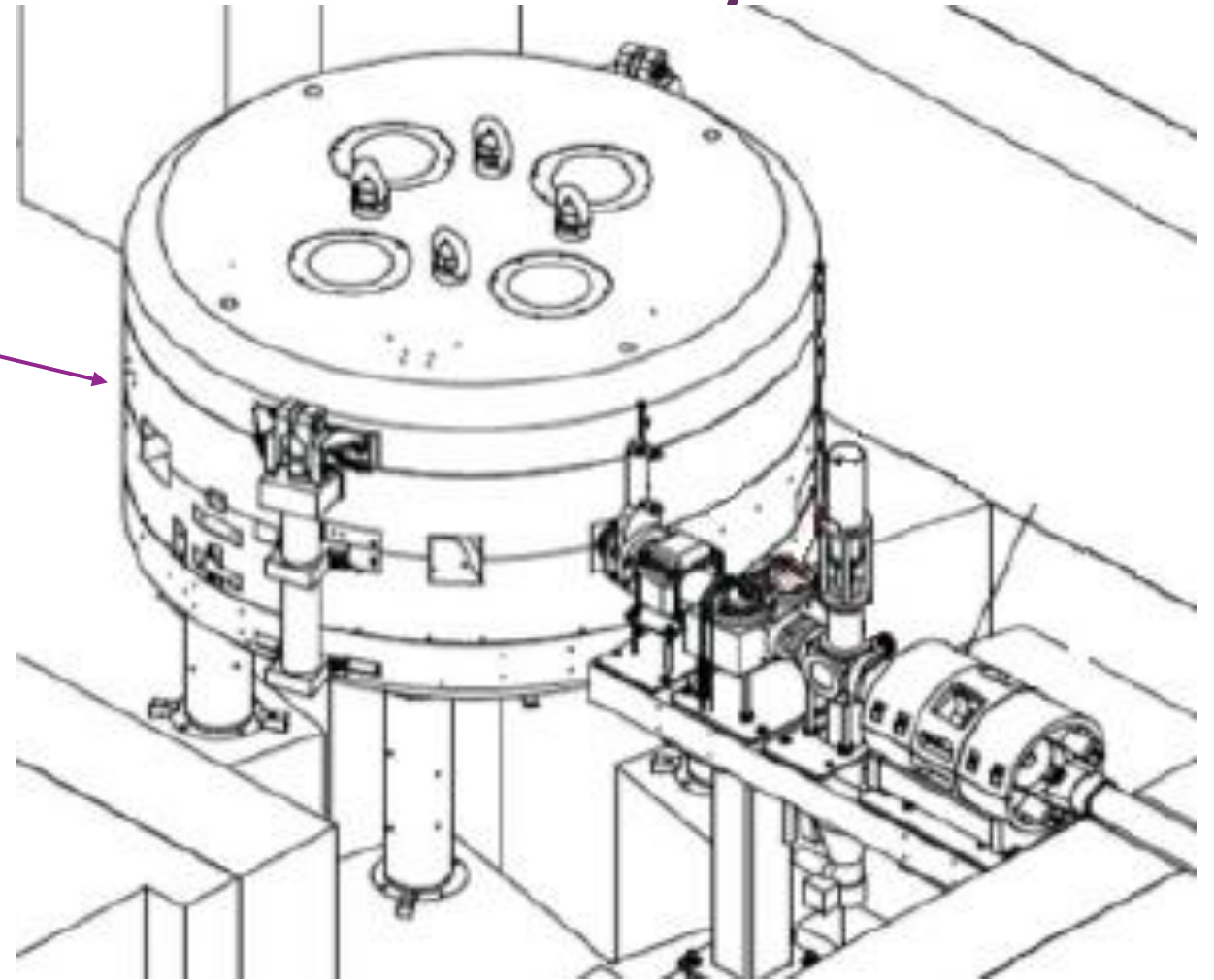
# The Bern Medical Cyclotron laboratory

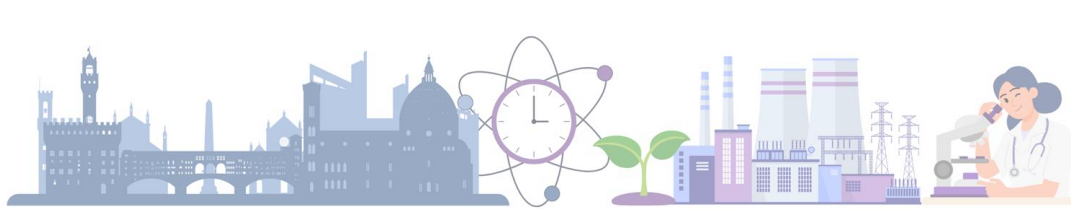
IBA 18/18 HC cyclotron



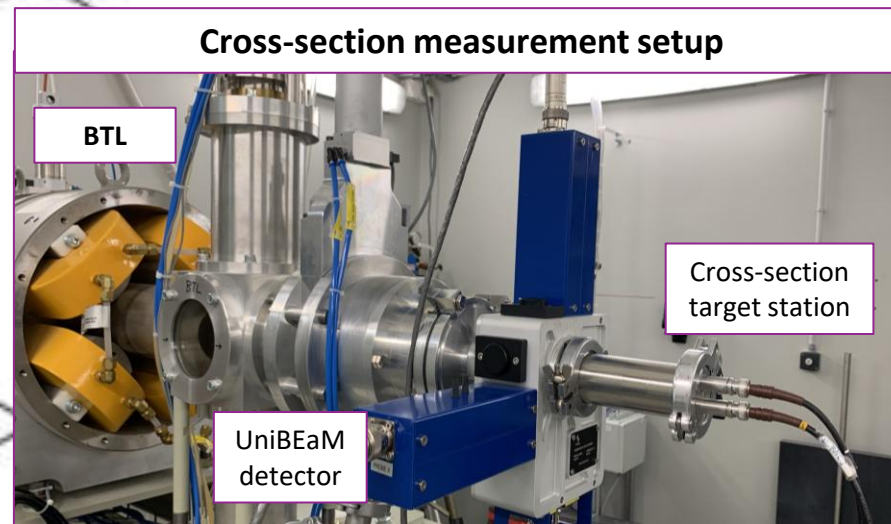
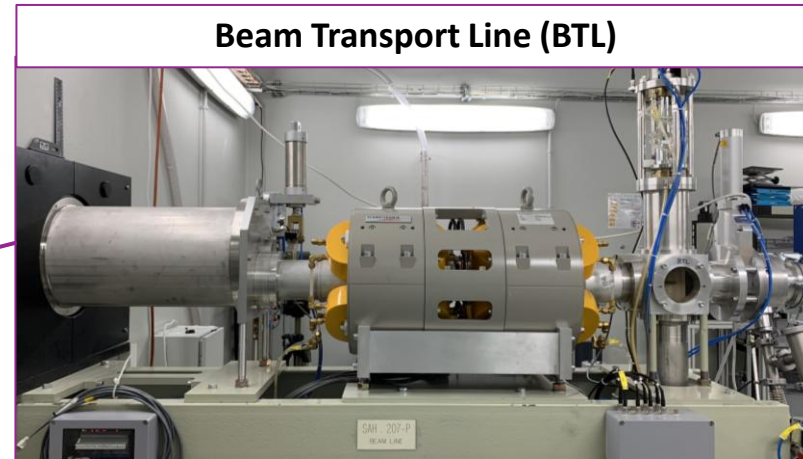
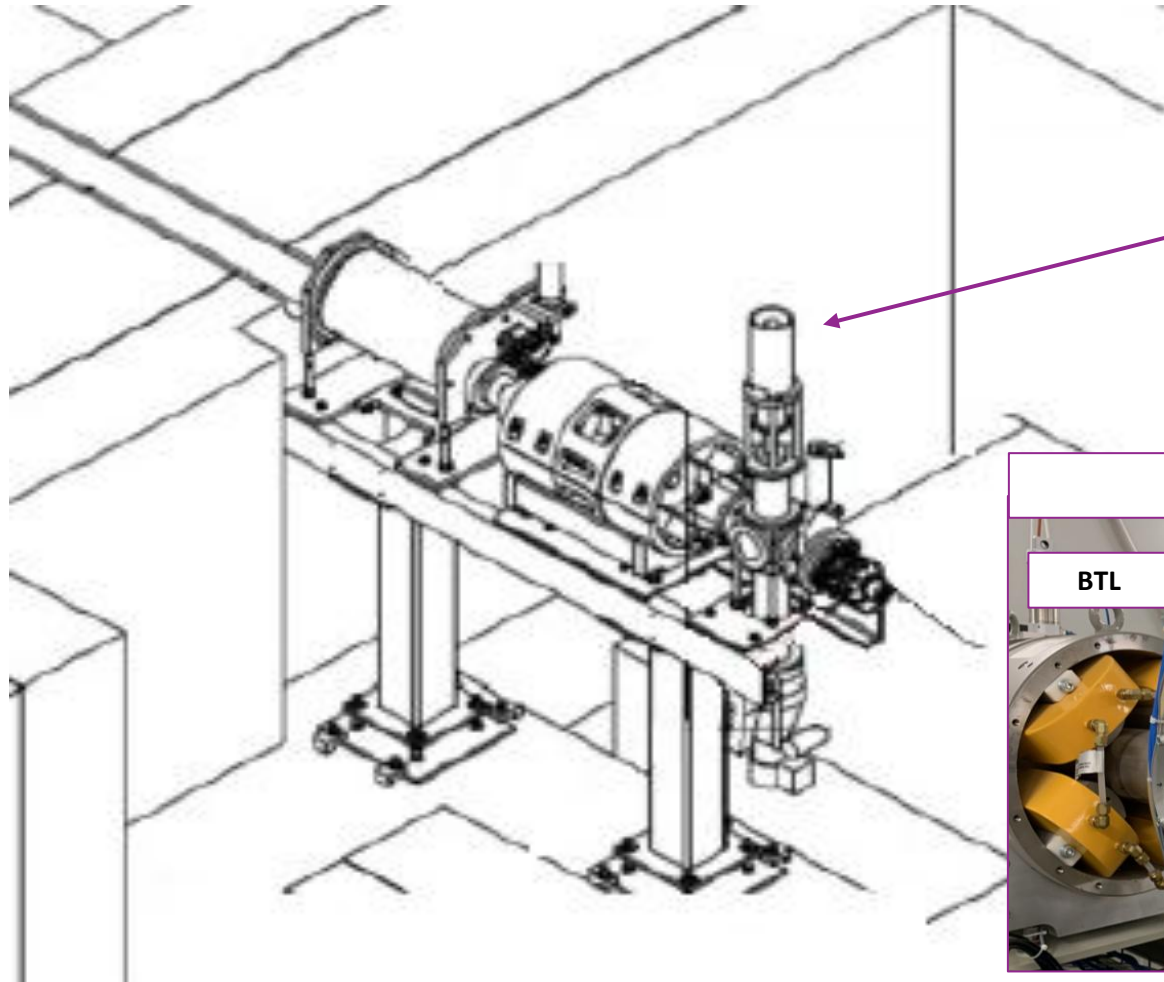
cyclotron  
configuration  
in 2023

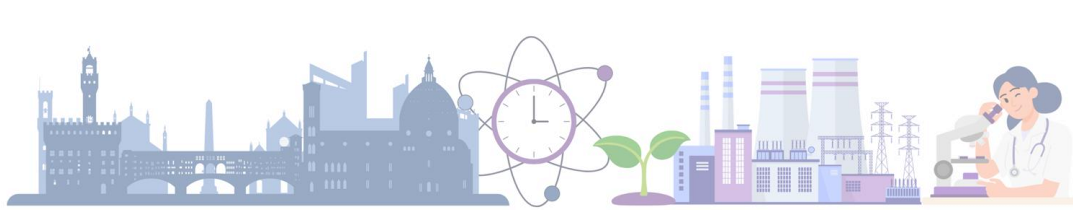
- Two H<sup>-</sup> ion sources
- High current (max 150 μA)
- 8 out ports:
  - 6 <sup>18</sup>F liquid targets [industrial production]
  - **Beam Transport Line (BTL) [research]**
  - **Solid Target Station (STS) [research]**



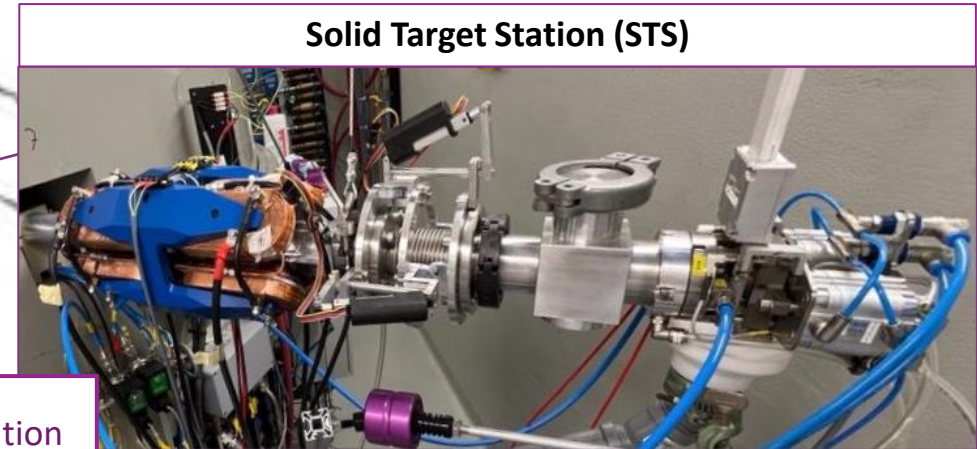
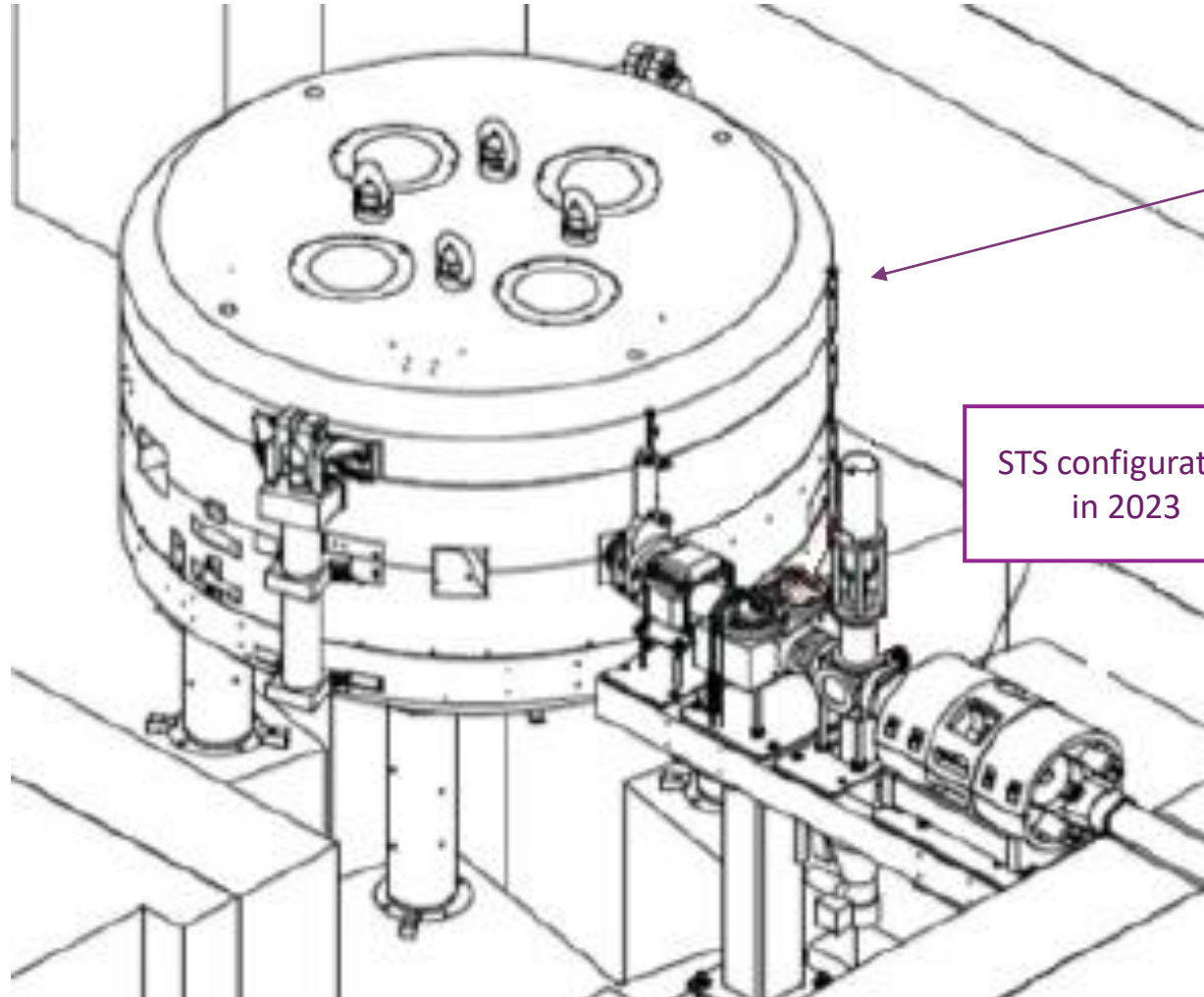


# The Bern Medical Cyclotron laboratory

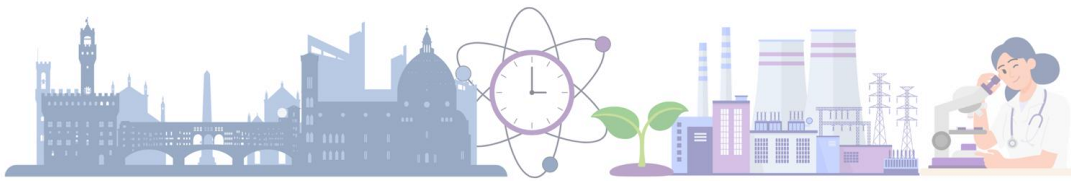




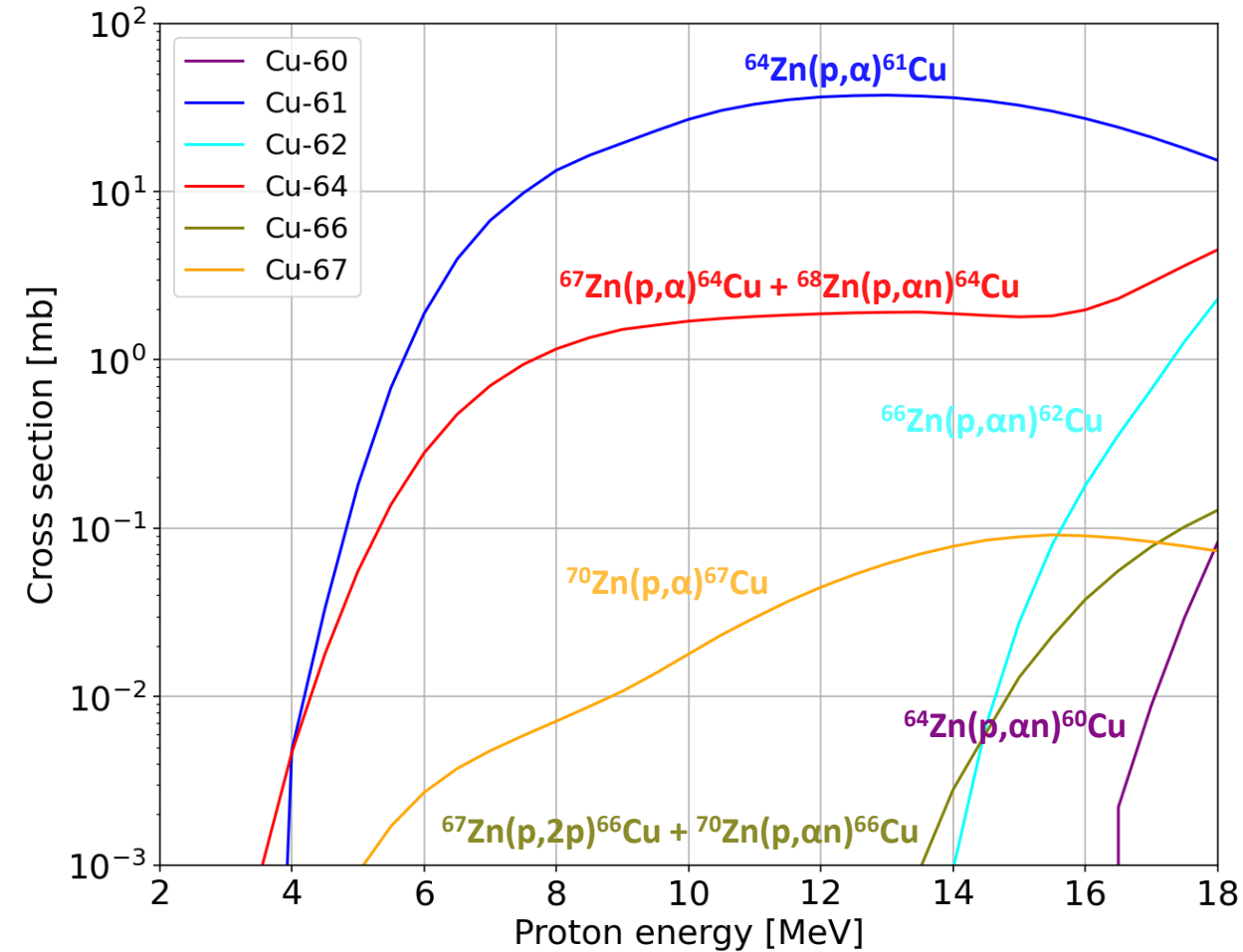
# The Bern Medical Cyclotron laboratory



STS configuration  
in 2023



# Proton irradiation of natural Zinc



	<sup>64</sup> Zn	<sup>66</sup> Zn	<sup>67</sup> Zn	<sup>68</sup> Zn	<sup>70</sup> Zn
Nat Zn [%]	49.17	27.73	4.04	18.45	0.61
Enr. <sup>64</sup> Zn [%]	<b>99.40</b>	0.39	0.04	0.17	<0.01
Enr. <sup>67</sup> ZnO [%]	1.56	3.88	<b>89.60</b>	4.91	0.05
Enr. <sup>68</sup> Zn [%]	0.99	0.81	0.38	<b>97.80</b>	0.02
Enr. <sup>70</sup> ZnO [%]	0.02	0.01	0.02	1.20	<b>98.75</b>

<sup>60</sup>Cu:  $t_{1/2} = 23.7$  (4) min

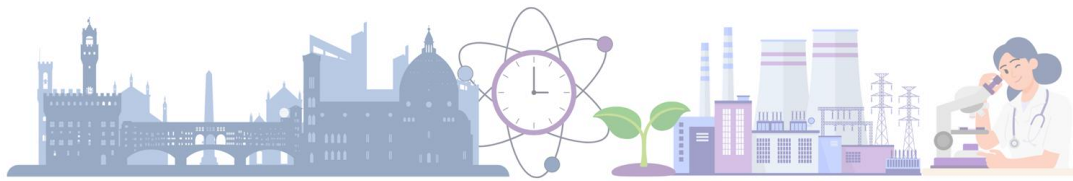
<sup>61</sup>Cu:  $t_{1/2} = 3.339$  (8) h

<sup>62</sup>Cu:  $t_{1/2} = 9.67$  (3) min

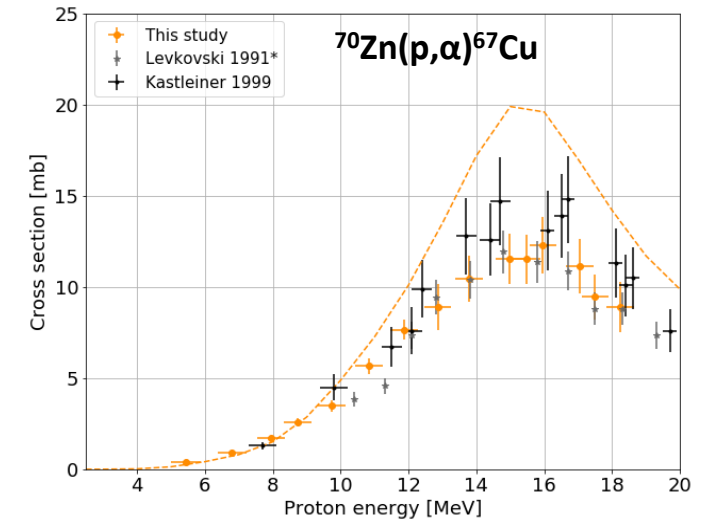
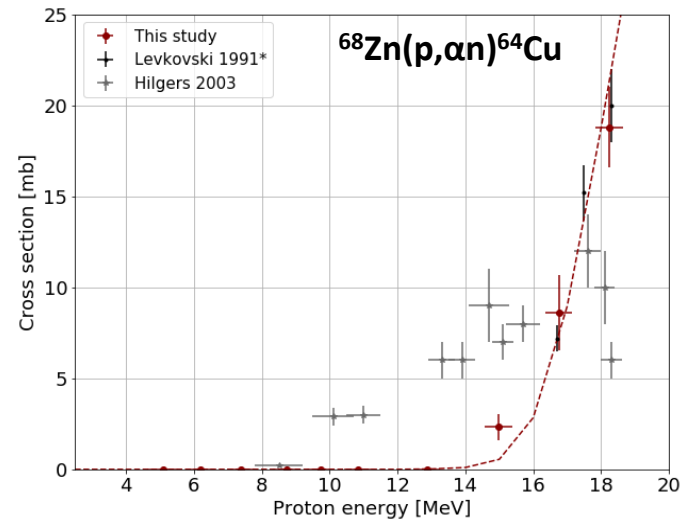
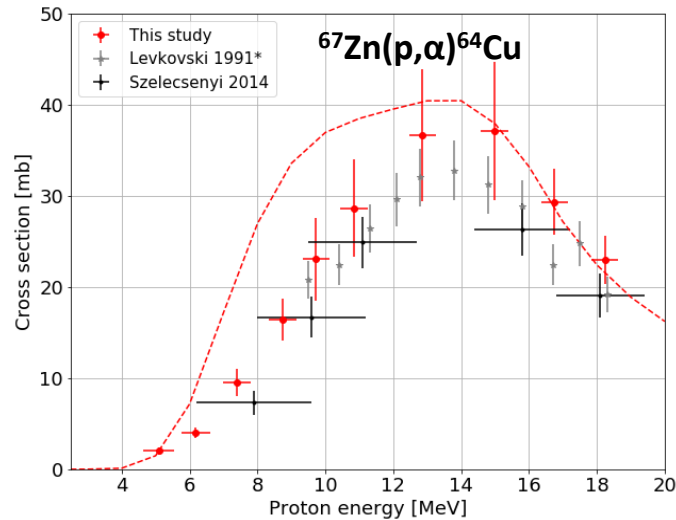
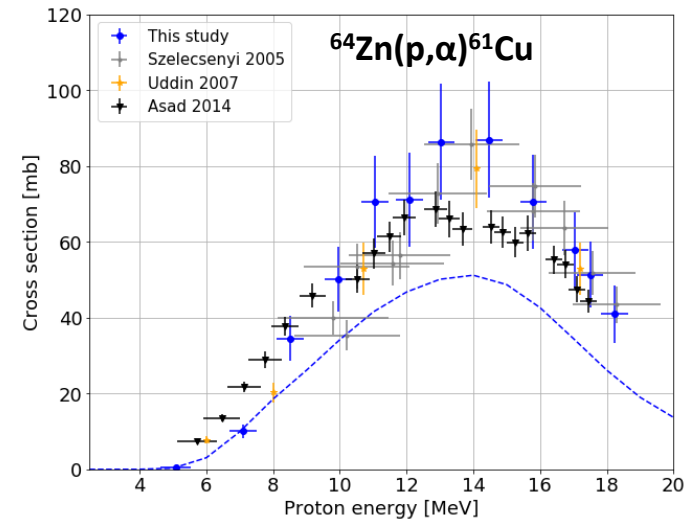
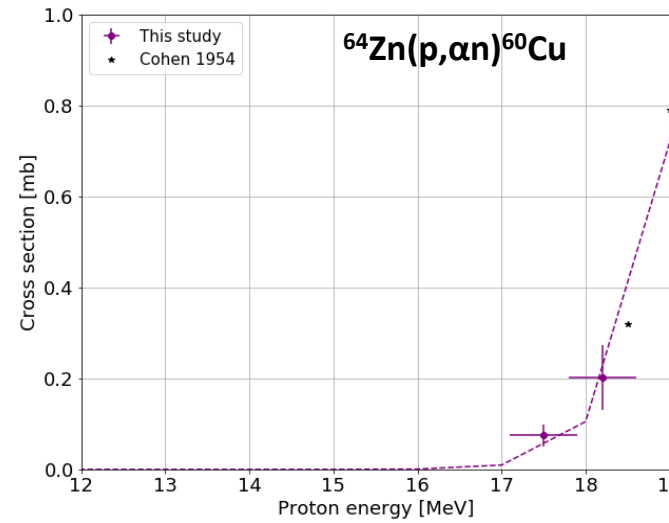
<sup>64</sup>Cu:  $t_{1/2} = 12.7006$  (20) h

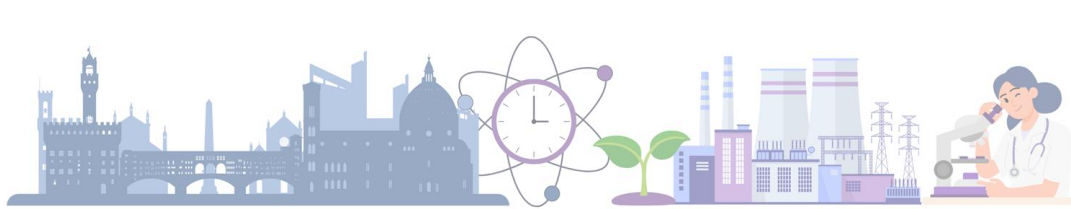
<sup>66</sup>Cu:  $t_{1/2} = 5.120$  (14) min

<sup>67</sup>Cu:  $t_{1/2} = 61.83$  (12) h



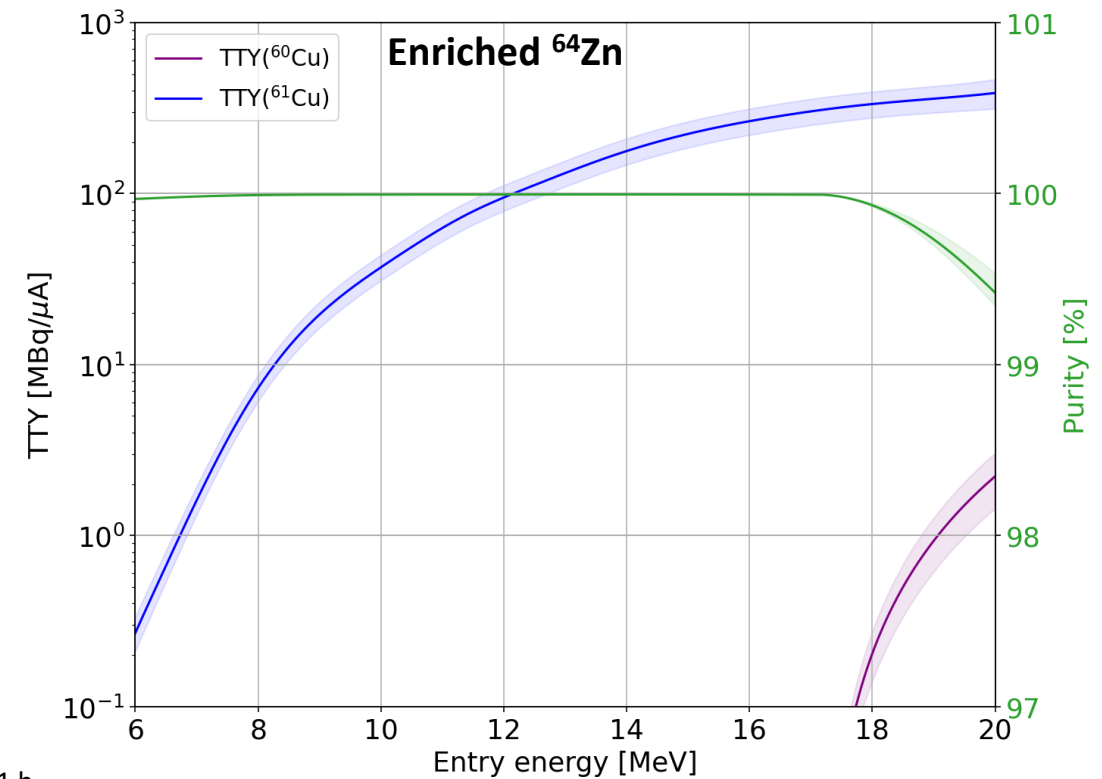
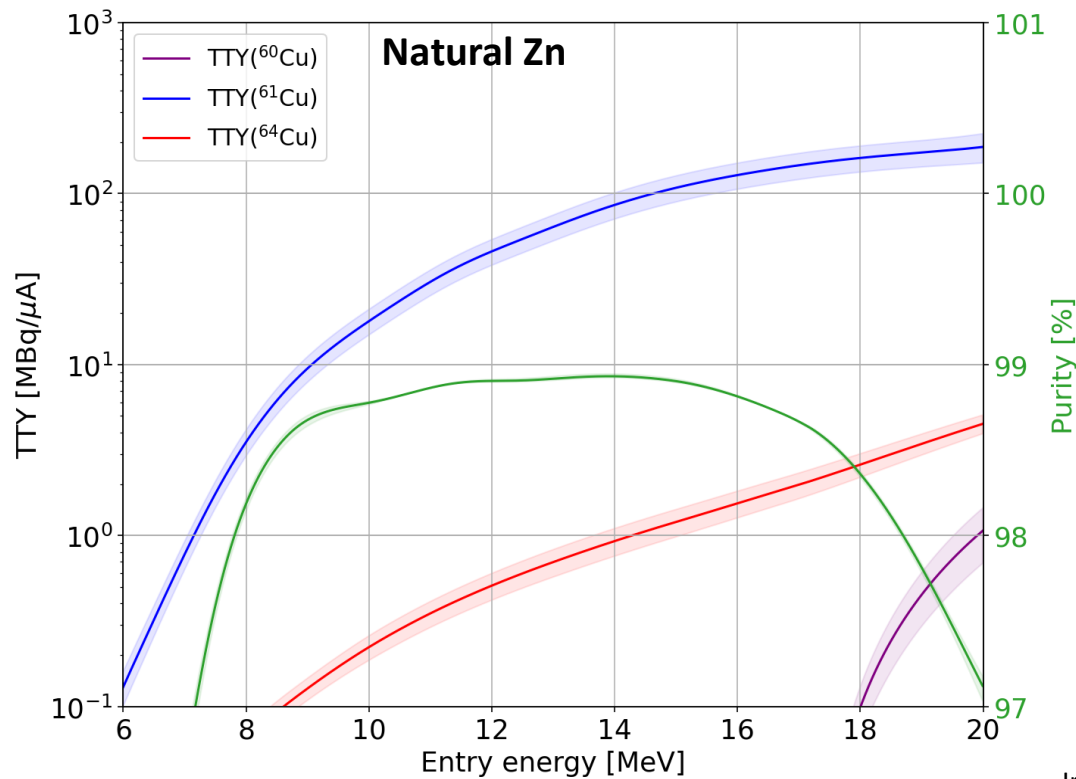
# Cross-section results



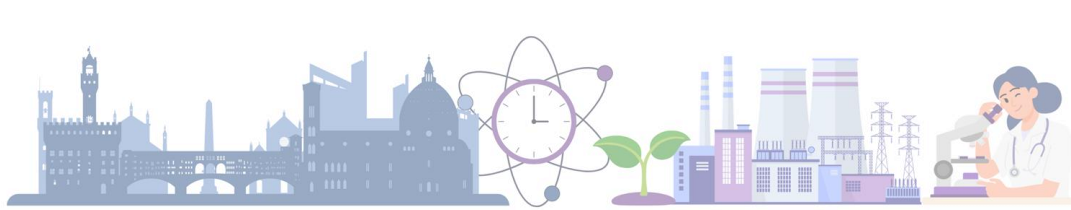


# $^{61}\text{Cu}$ production: natural vs enriched $^{64}\text{Zn}$

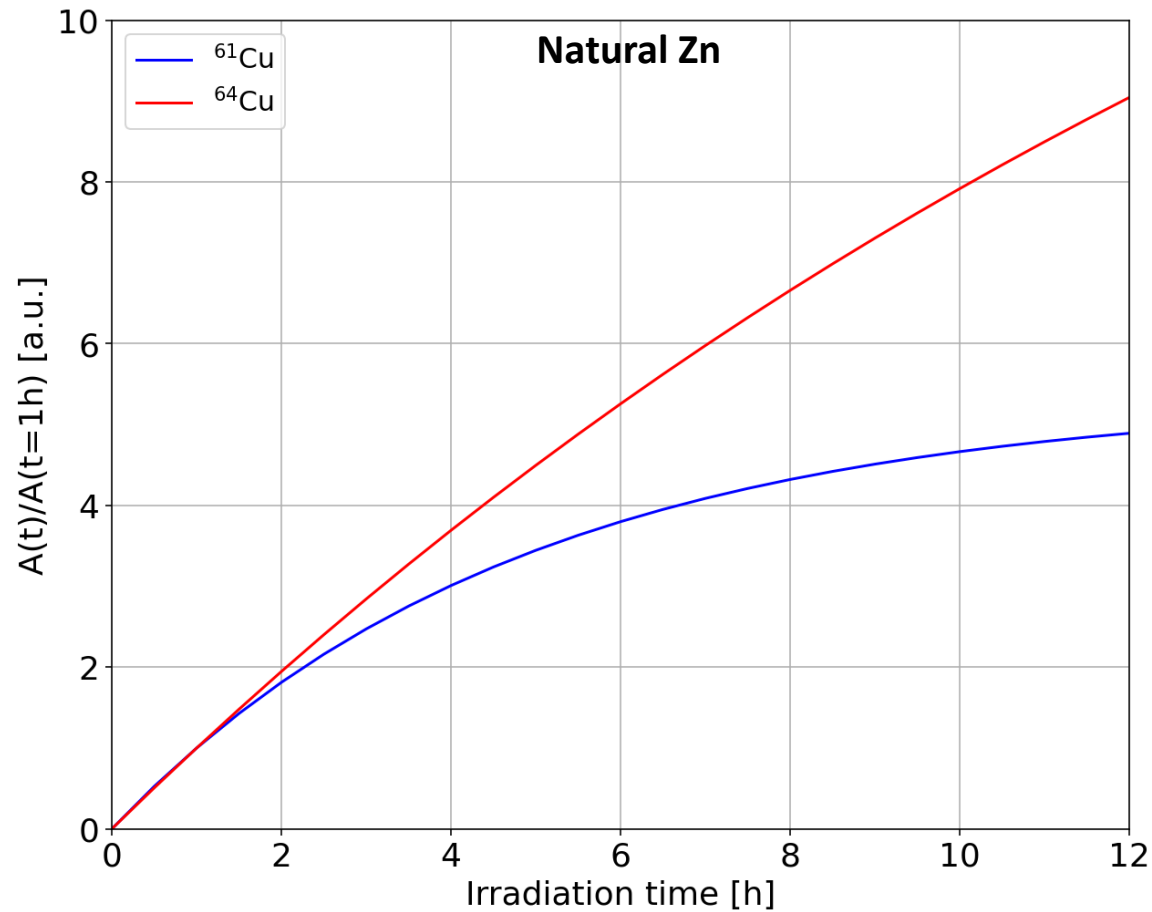
	$^{64}\text{Zn}$	$^{66}\text{Zn}$	$^{67}\text{Zn}$	$^{68}\text{Zn}$	$^{70}\text{Zn}$
Nat Zn [%]	49.17	27.73	4.04	18.45	0.61
Enr. $^{64}\text{Zn}$ [%]	99.40	0.39	0.04	0.17	<0.01

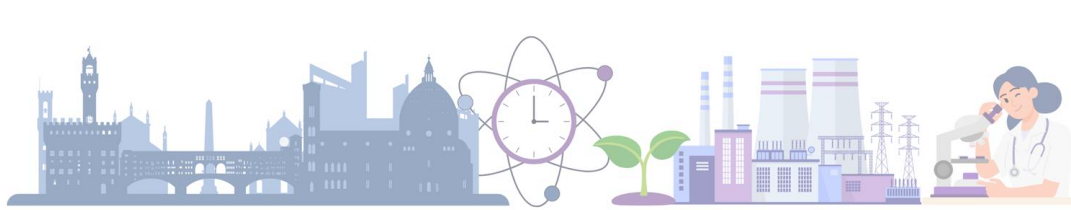


Irradiation time = 1 h



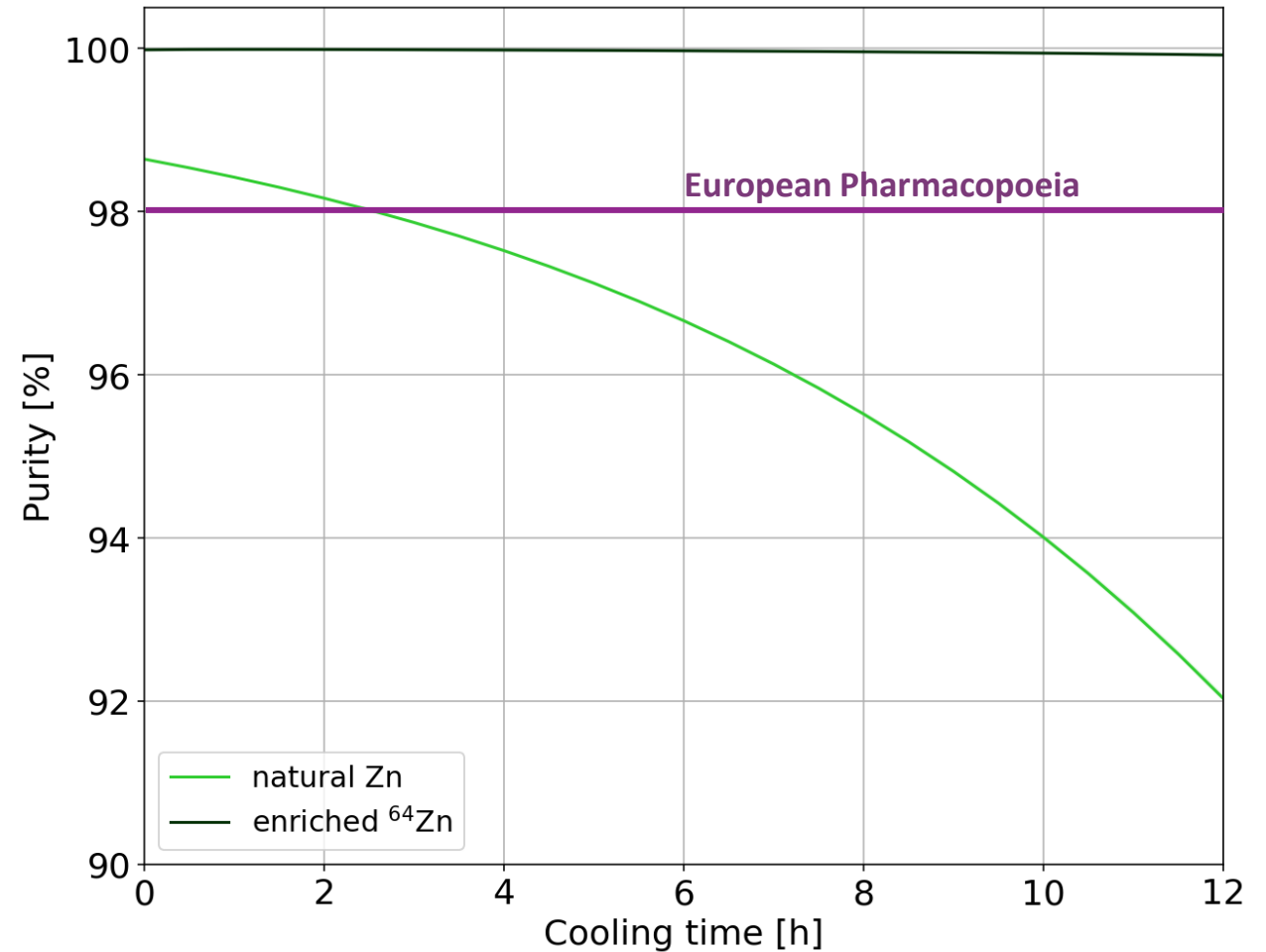
# $^{61}\text{Cu}$ production: natural vs enriched $^{64}\text{Zn}$

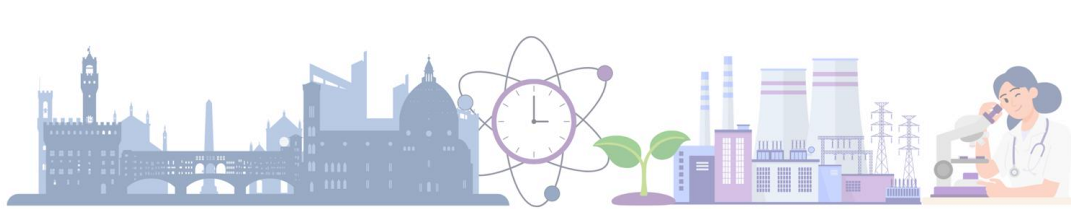




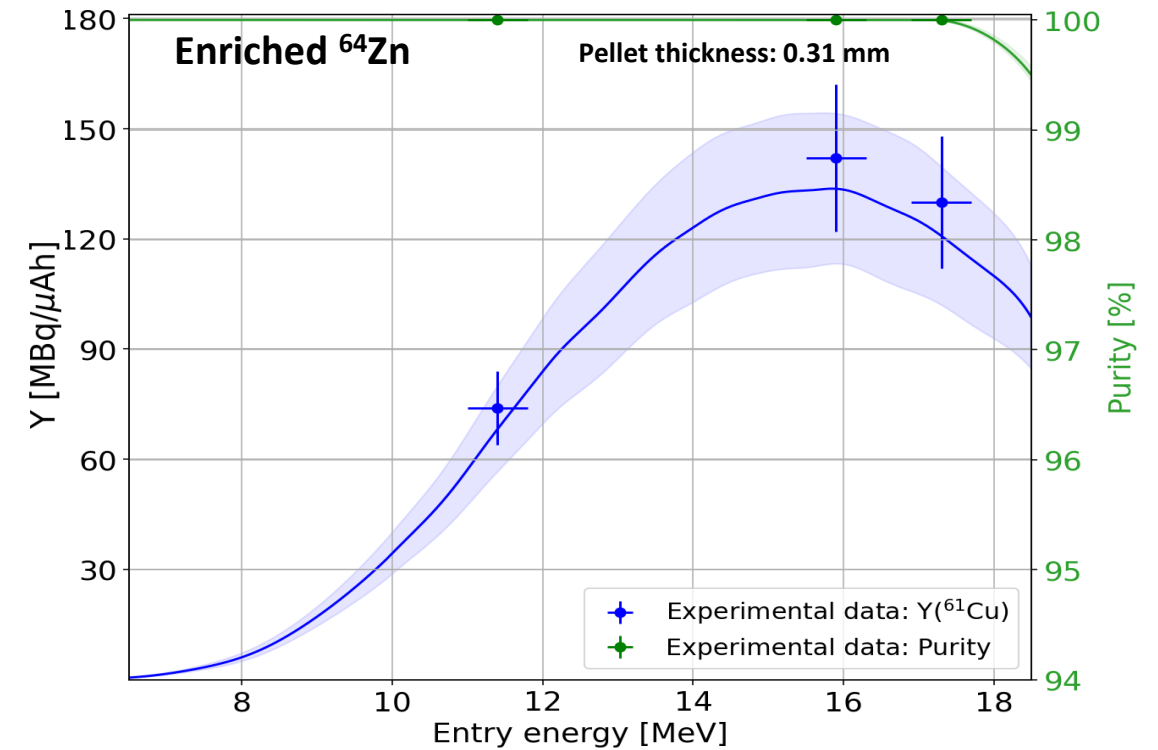
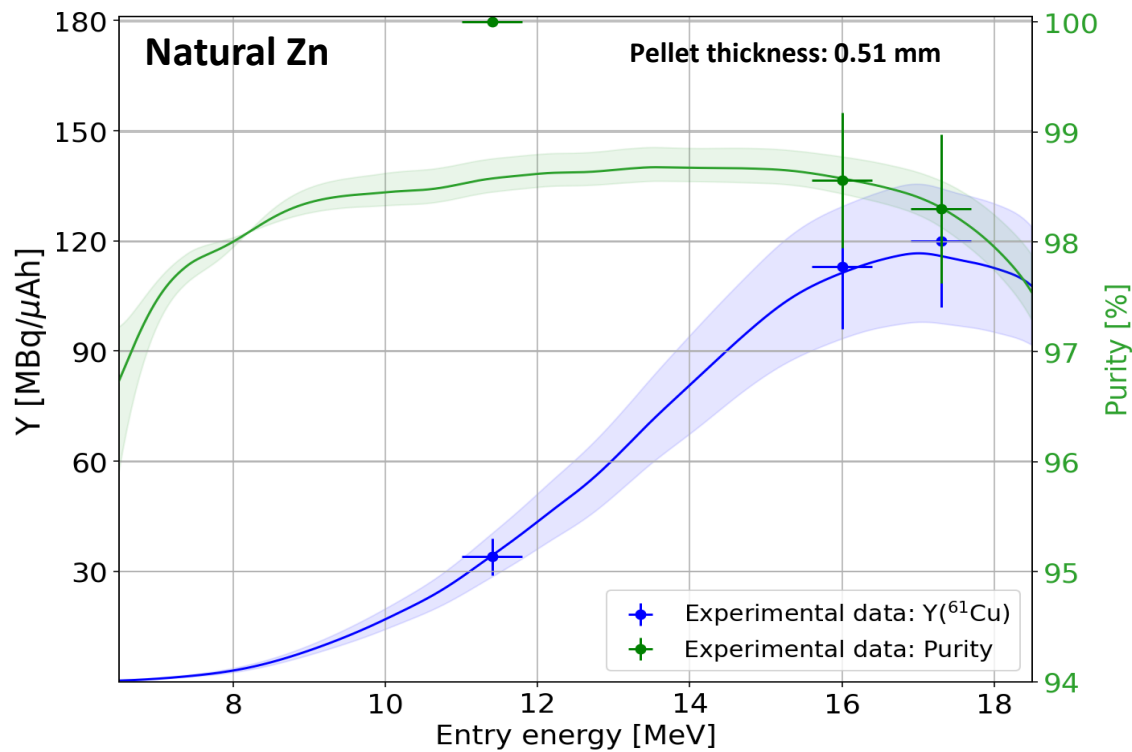
# $^{61}\text{Cu}$ production: natural vs enriched $^{64}\text{Zn}$

	Nat Zn	Enr $^{64}\text{Zn}$
E [MeV]	16.0	17.8
Irrad time [h]	3	12
TTY [MBq/ $\mu\text{A}$ ]	317	1606
$P_{\text{EOB}}$ [%]	98.64	99.98



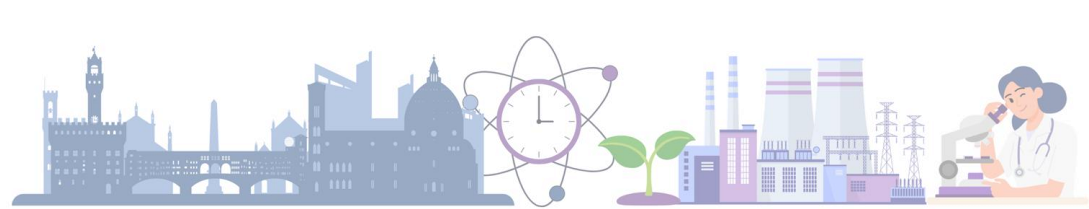


# <sup>61</sup>Cu production tests



	E [MeV]	Q [10 <sup>-4</sup> μAh]	Y( <sup>61</sup> Cu) [MBq/μAh]	P( <sup>61</sup> Cu) [%]	Q [10 <sup>-4</sup> μAh]	Y( <sup>61</sup> Cu) [MBq/μAh]	P( <sup>61</sup> Cu) [%]
<b>Test 1</b>	17.3 ± 0.4	3.3 ± 0.3	120 ± 18 (115)	98.3 ± 0.7 (98.3)	2.3 ± 0.2	130 ± 18 (123)	100 (100)
<b>Test 2</b>	16.0 ± 0.4	3.1 ± 0.3	113 ± 17 (111)	98.6 ± 0.6 (98.6)	2.7 ± 0.3	142 ± 20 (131)	100 (100)
<b>Test 3</b>	11.4 ± 0.4	1.1 ± 0.1	34 ± 5 (35)	100 (99)	1.1 ± 0.1	74 ± 10 (70)	100 (100)

G. Dellepiane et al, *Appl Radiat Isot* 2022, 190, 110466

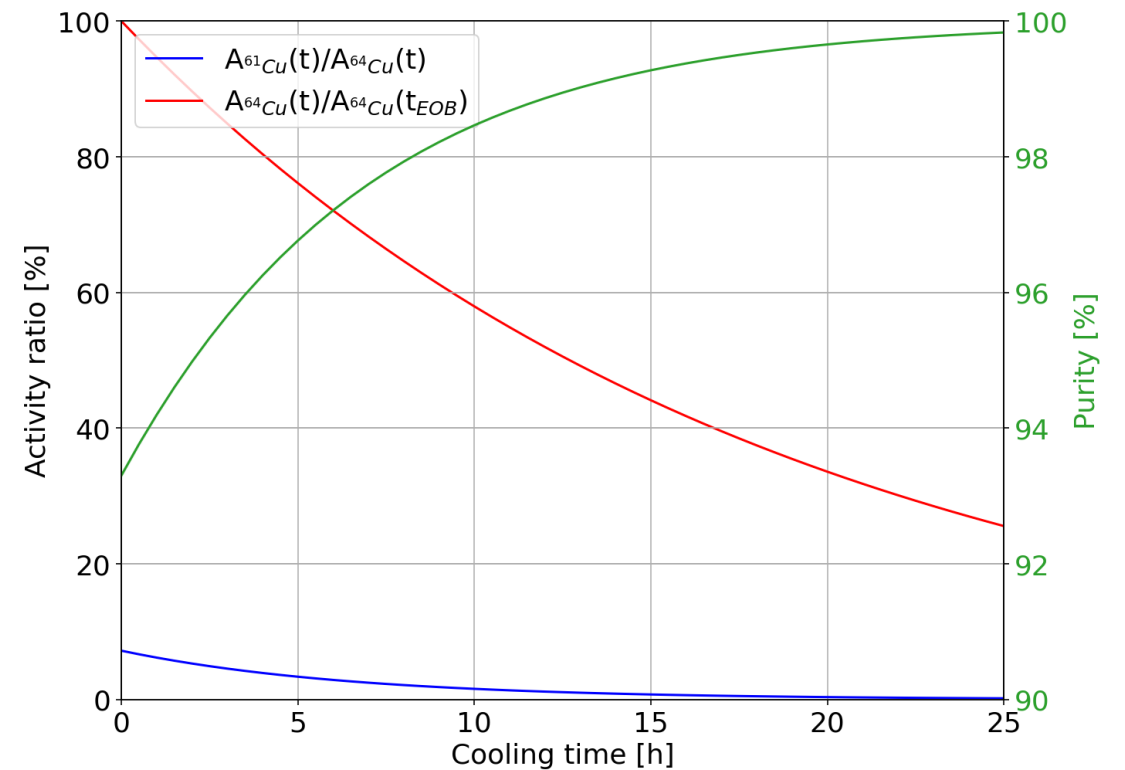
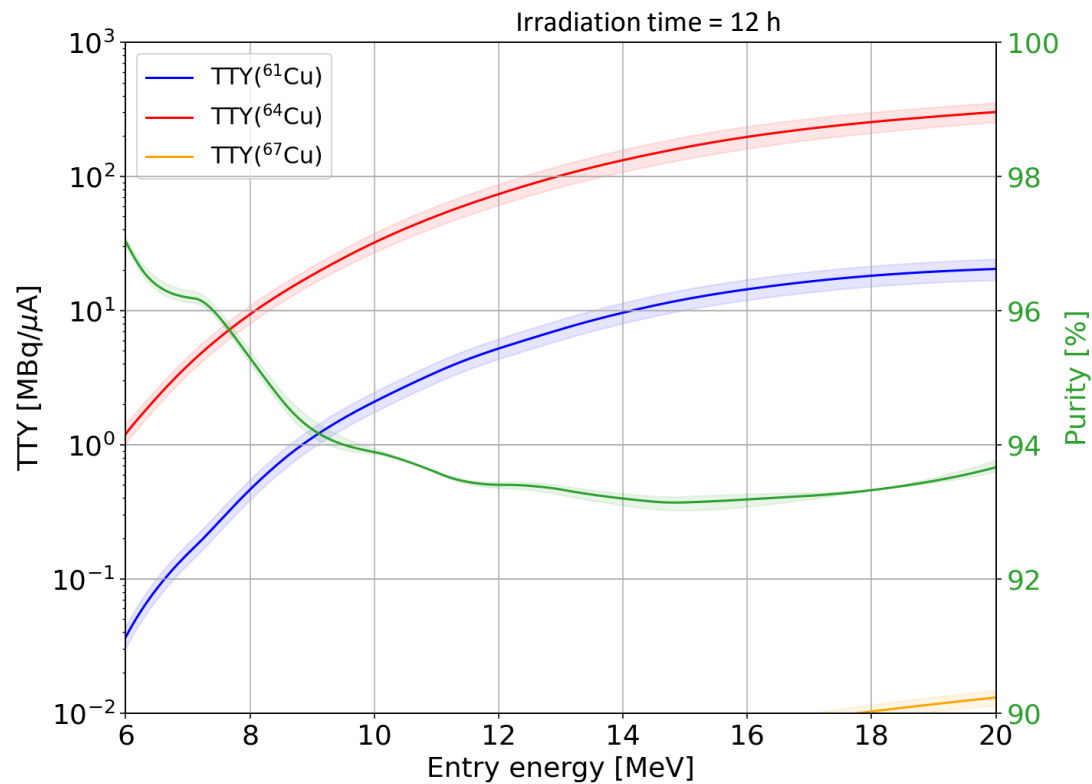


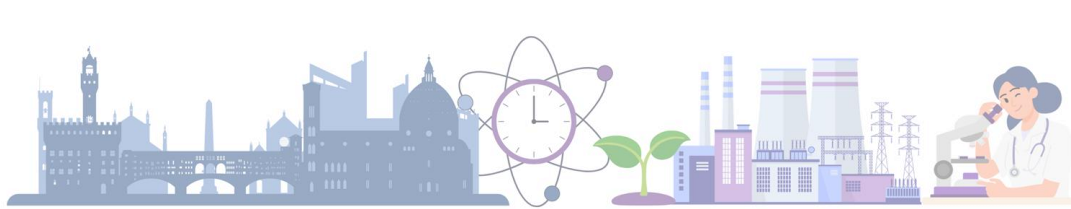
# $^{64}\text{Cu}$ production

	$^{64}\text{Zn}$	$^{66}\text{Zn}$	$^{67}\text{Zn}$	$^{68}\text{Zn}$	$^{70}\text{Zn}$
Enr. $^{67}\text{ZnO}$ [%]	1.56	3.88	<b>89.60</b>	4.91	0.05

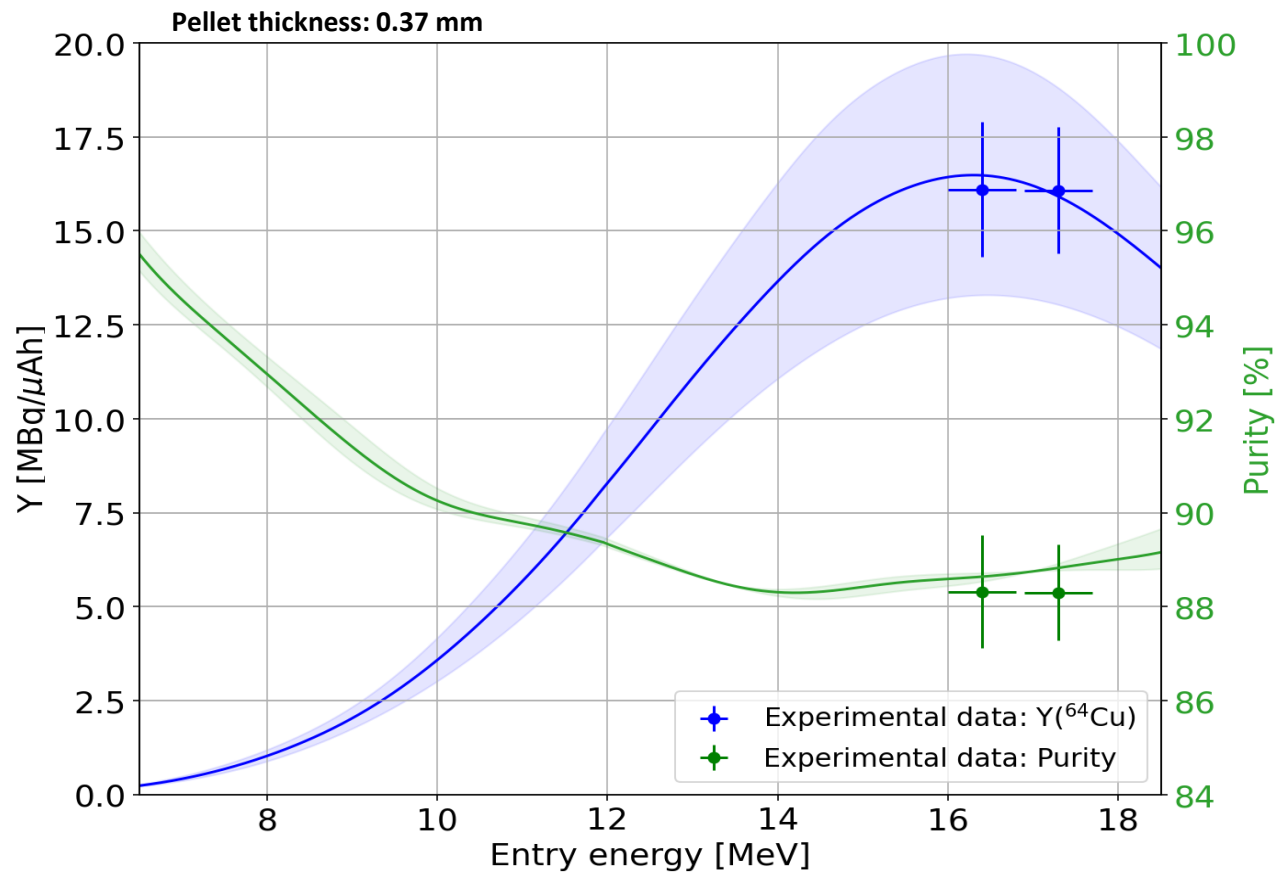
	EOB	$t_c = 10$ h
TTY [MBq/ $\mu\text{A}$ ]	250	170
$P_{\text{EOB}}$ [%]	93.30	98.47

\*Proton energy: **17.8 MeV**; Irradiation time: **12 h**



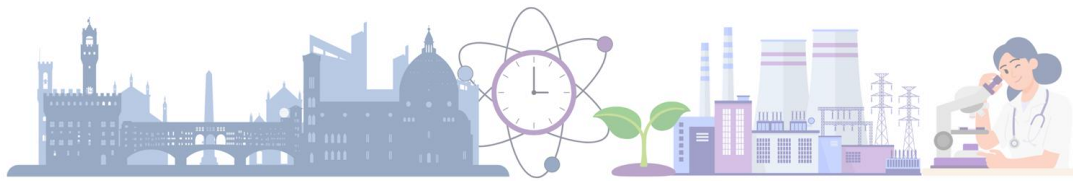


# <sup>64</sup>Cu production tests



	E [MeV]	Q [10 <sup>-4</sup> μAh]	Y( <sup>64</sup> Cu) [MBq/μAh]	P <sub>EOB</sub> ( <sup>64</sup> Cu) [%]	P <sub>10h</sub> ( <sup>64</sup> Cu) [%]
<b>Test 1</b>	17.3 ± 0.4	2.7 ± 0.3	16.1 ± 1.7 (15.9)	88.3 ± 1.0 (88.8)	97.4 ± 1.0 (97.4)
<b>Test 2</b>	16.4 ± 0.4	1.3 ± 0.1	16.1 ± 1.8 (16.3)	88.3 ± 1.2 (88.6)	97.3 ± 1.3 (97.3)

G. Dellepiane et al, *Appl Radiat Isot* 2023, 191, 110518

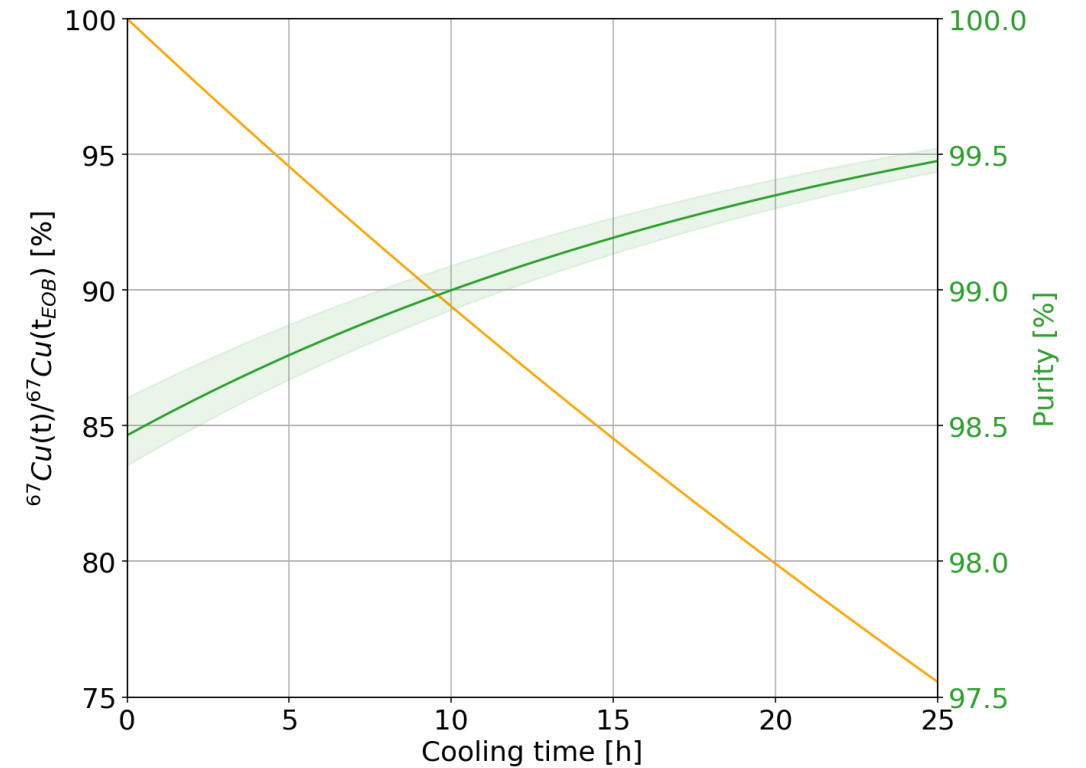
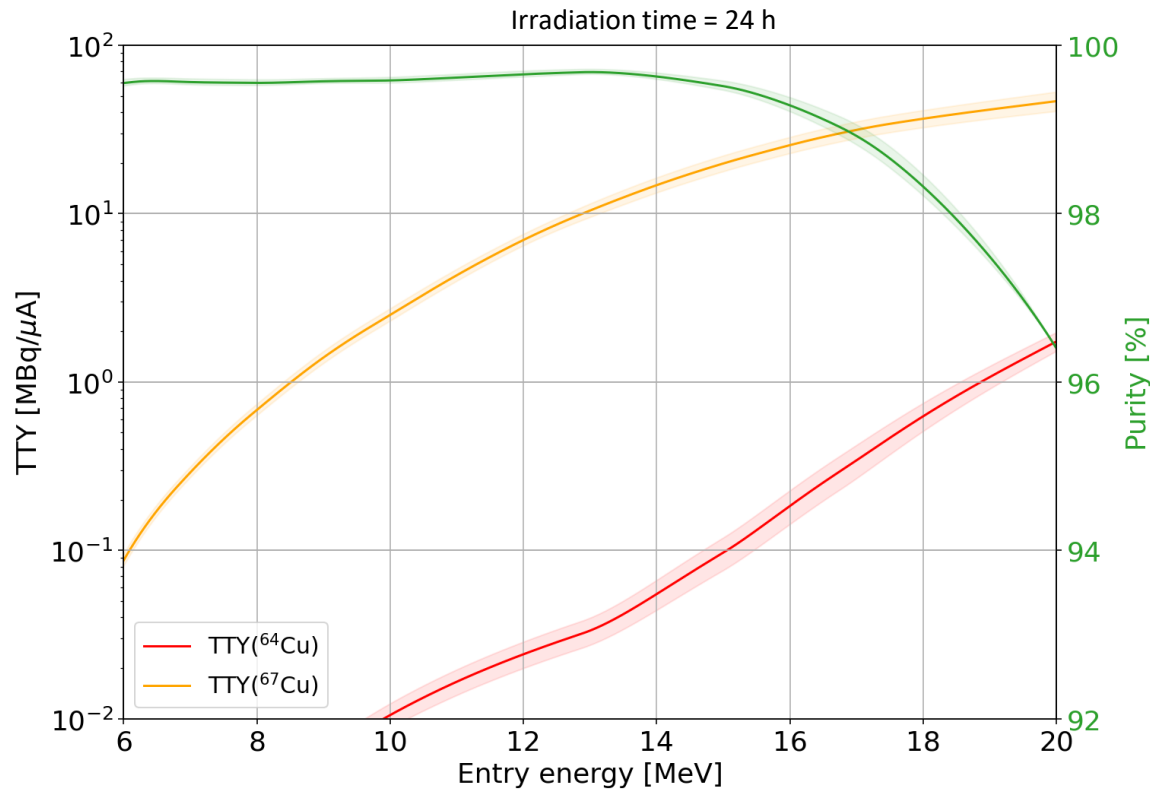


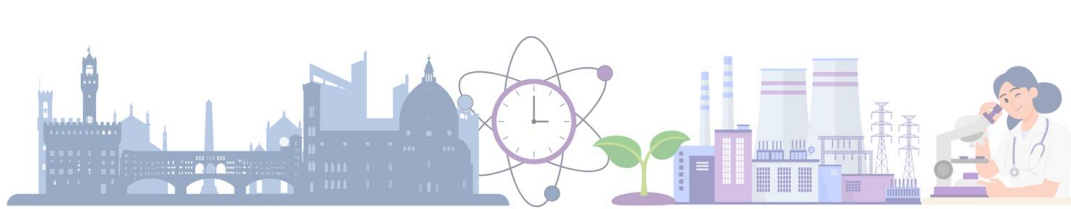
# <sup>67</sup>Cu production

	<sup>64</sup> Zn	<sup>66</sup> Zn	<sup>67</sup> Zn	<sup>68</sup> Zn	<sup>70</sup> Zn
Enr. <sup>70</sup> ZnO [%]	0.02	0.01	0.02	1.20	<b>98.75</b>

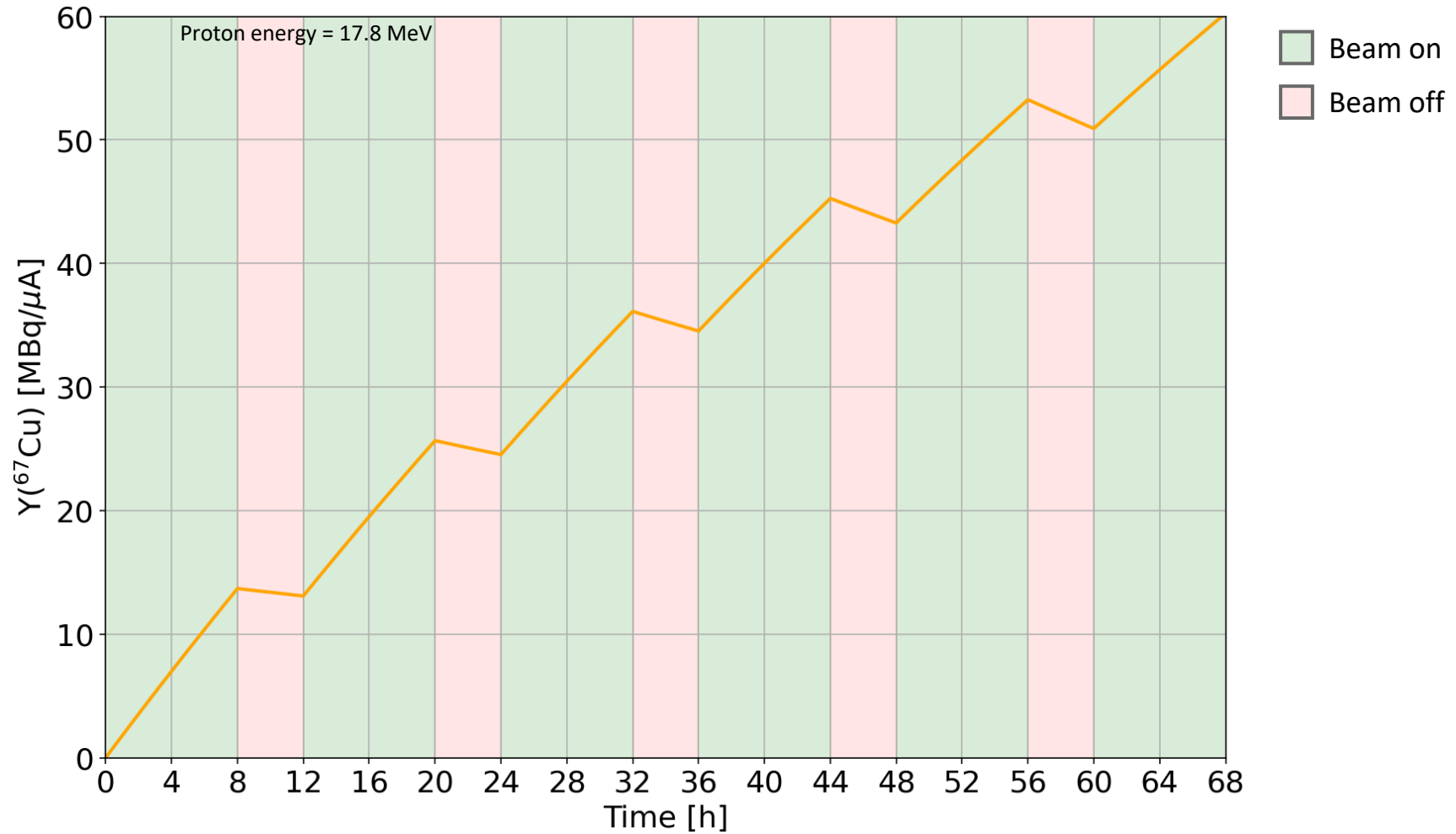
	EOB	t <sub>c</sub> = 10 h
TTY [MBq/μA]	36	31
P <sub>EOB</sub> [%]	98.46	99.00

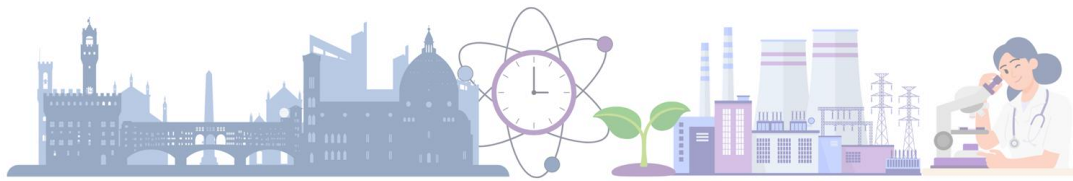
\*Proton energy: **17.8 MeV**; Irradiation time: **24 h**



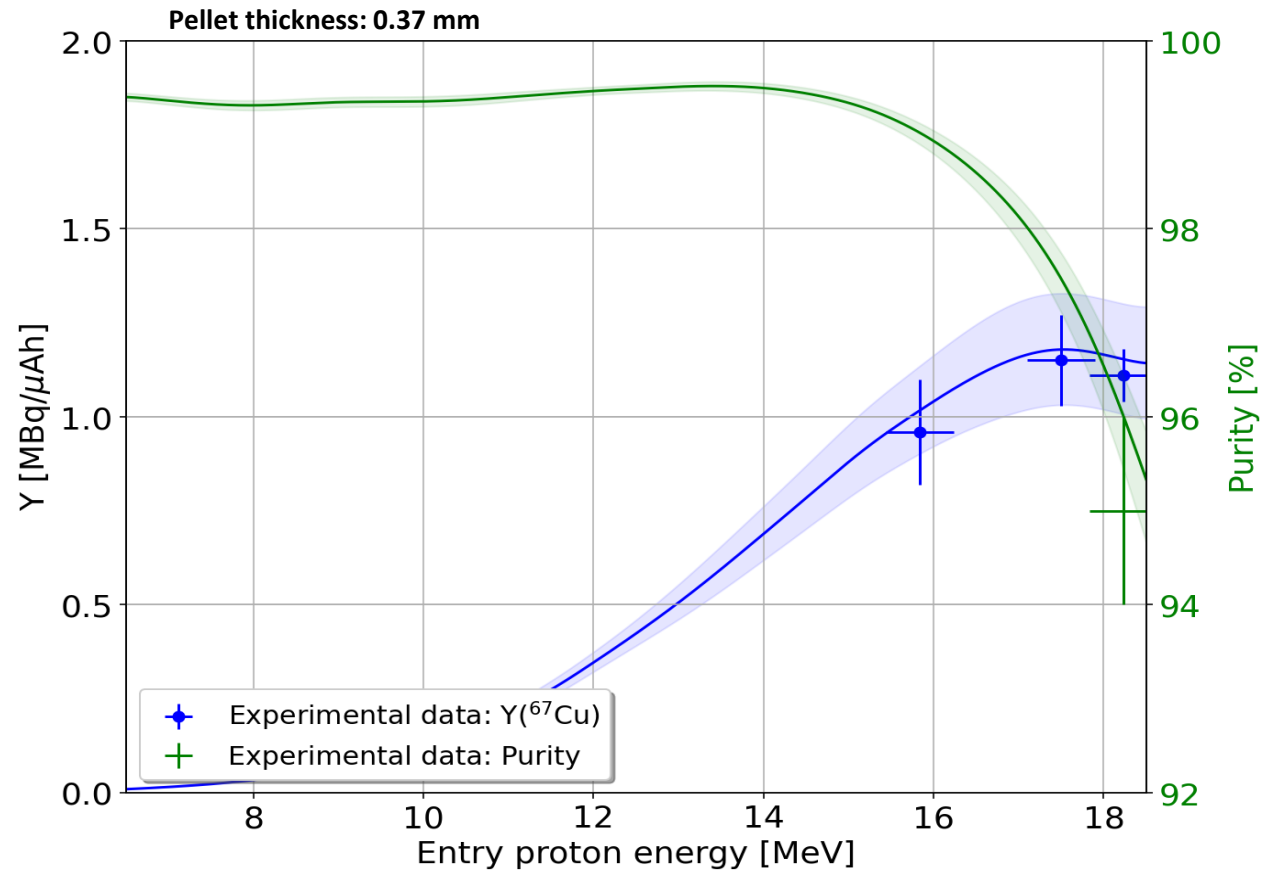


# $^{67}\text{Cu}$ production



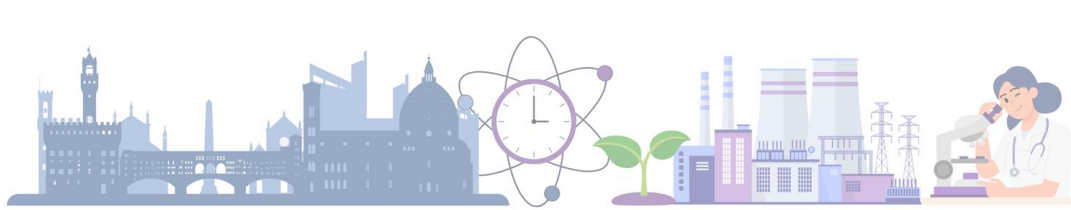


# <sup>67</sup>Cu production tests

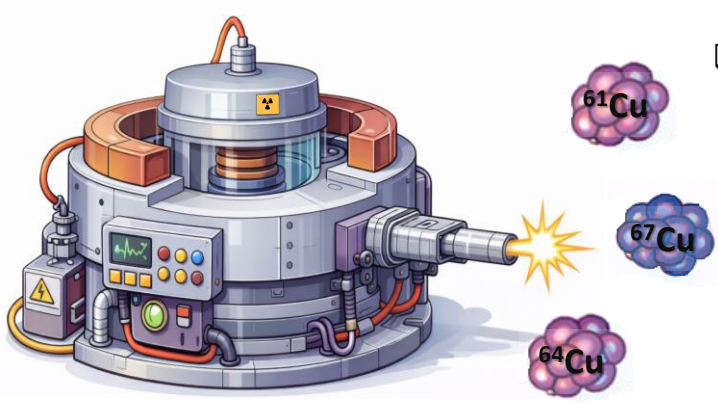


	E [MeV]	Q [ $10^{-3}$ μAh]	$Y(^{64}\text{Cu})$ [MBq/μAh]	$P_{\text{EOB}}(^{64}\text{Cu})$ [%]
<b>Test 1</b>	$11.0 \pm 0.4$	$7.9 \pm 0.8$	$0.96 \pm 0.14$ (1.02)	100 (99)
<b>Test 2</b>	$13.2 \pm 0.4$	$1.7 \pm 0.2$	$1.15 \pm 0.12$ (1.18)	100 (98)
<b>Test 3</b>	$14.0 \pm 0.4$	$37.7 \pm 1.8$	$1.11 \pm 0.07$ (1.15)	$95 \pm 1$ (96)

G. Dellepiane et al, *Appl Radiat Isot* 2023, 195, 110737

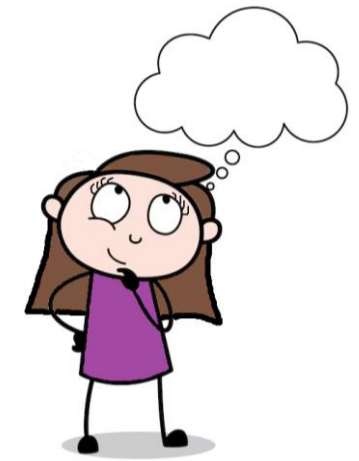


# Conclusions and outlook



- ❑ We demonstrated the feasibility of producing  $^{61}\text{Cu}$ ,  $^{64}\text{Cu}$  and  $^{67}\text{Cu}$  using a biomedical cyclotron
- ❑ **Highly-enriched zinc targets** are essential to achieve **high radionuclidic purity**  
**ALTHOUGH low-cost natural Zn** may be considered for nearby  $^{61}\text{Cu}$  PET applications

- ❑ For  $^{61}\text{Cu}$  and  $^{64}\text{Cu}$ , the achievable yields are sufficient for pre-clinical and clinical applications
- ❑  $^{67}\text{Cu}$  production requires **high beam currents**  
**THEREFORE further studies** are needed to assess **target robustness** and **thermal performance** under high-current irradiations





**Thank you all for your attention!**

