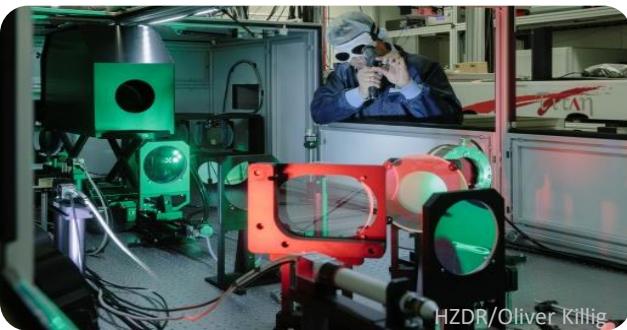
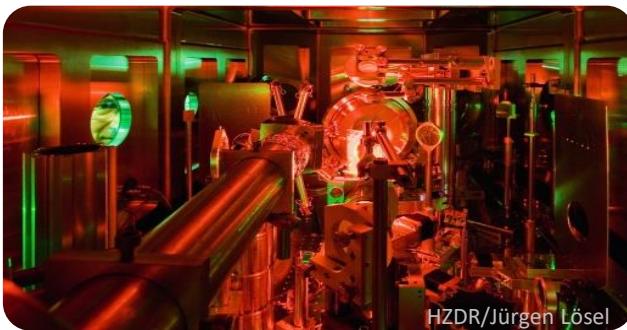


High energy proton acceleration at DRACO-PW and radio-biological applications



HZDR/Oliver Killig



HZDR/Jürgen Lösel



HZDR/Oliver Killig



HZDR/Detlev Müller

Karl Zeil

Helmholtz-Zentrum Dresden-Rossendorf

20th Advanced Accelerator Concepts Workshop (AAC'22)

November 7, 2022

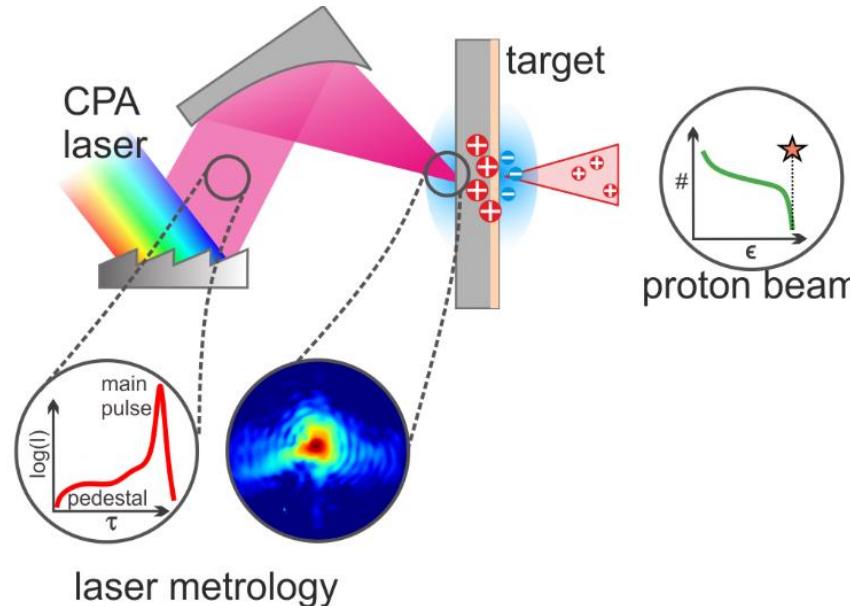
hzdr

M T ARD

ATHENA
The Helmholtz-Project for
Laser-Plasma-Acceleratio

Oncoray®
Regional Center for
Radiation Research in Oncology
Dresden

Development of a high repetition-rate, high energy plasma accelerator for ions

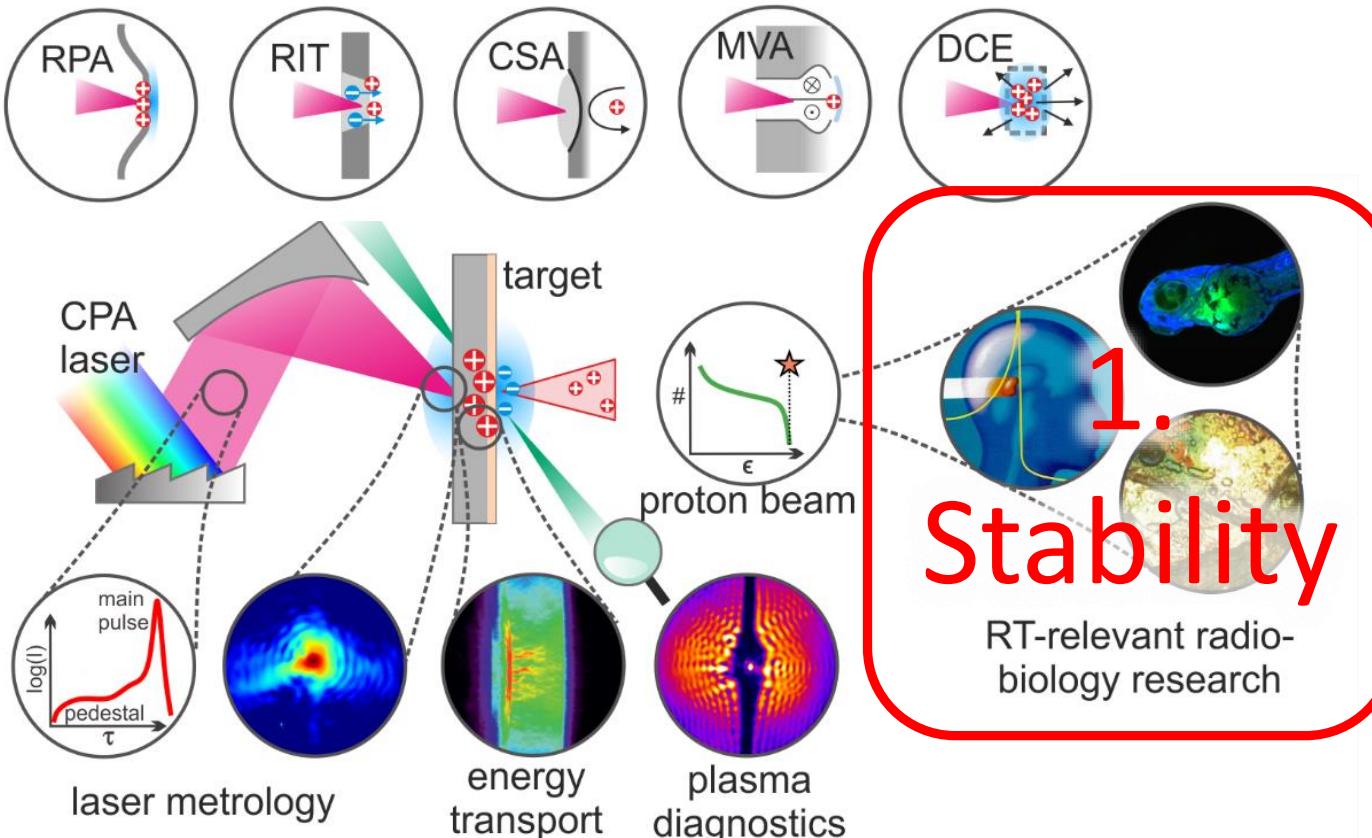


Energy scaling – two challenges:

1. Technological limits for larger laser systems:
 - Advanced accelerator schemes: from surface to volumetric interaction
 - Indirect, highly non-linear processes (instabilities) → high sensitivity on input parameters

2020 Roadmap on plasma accelerators
New Journal of Physics 23, 031101 (2021)

Development of a high repetition-rate, high energy plasma accelerator for ions



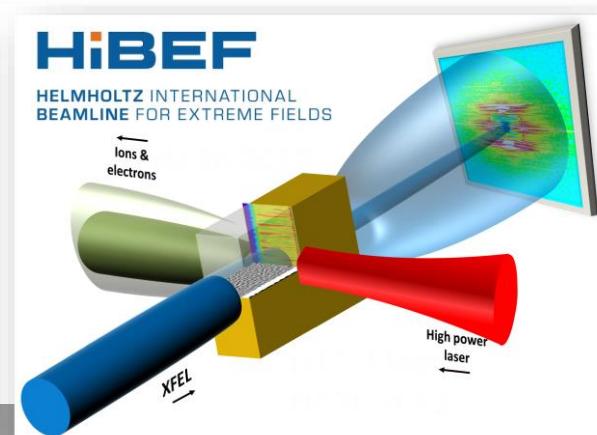
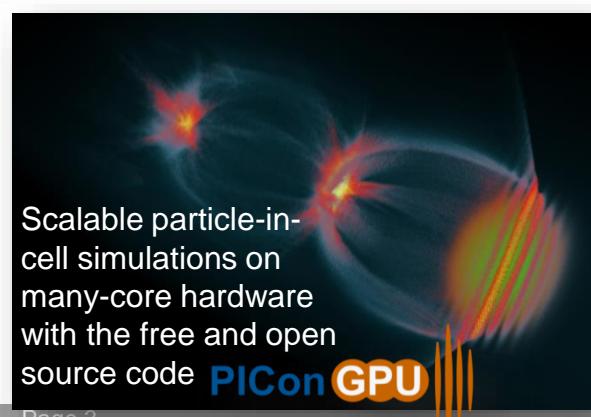
Energy scaling – two challenges:

1. Technological limits for larger laser systems:

- Advanced accelerator schemes: from surface to volumetric interaction
- Indirect, highly non-linear processes (instabilities) → high sensitivity on input parameters

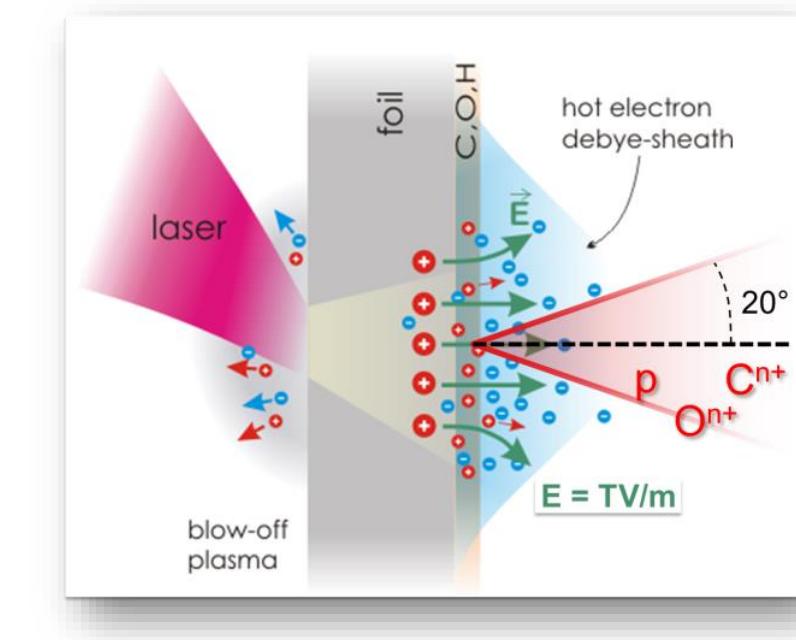
2. Limited predictability of simulations

- microscopic understanding
- control and knowledge of laser parameters on target

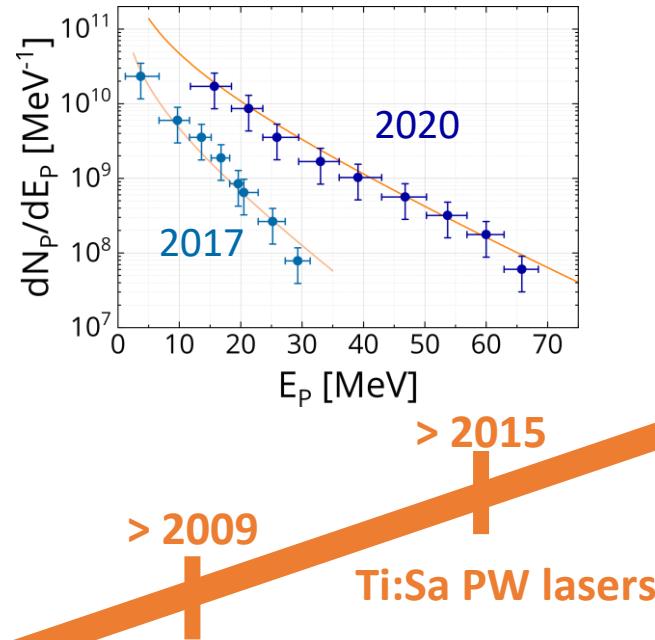
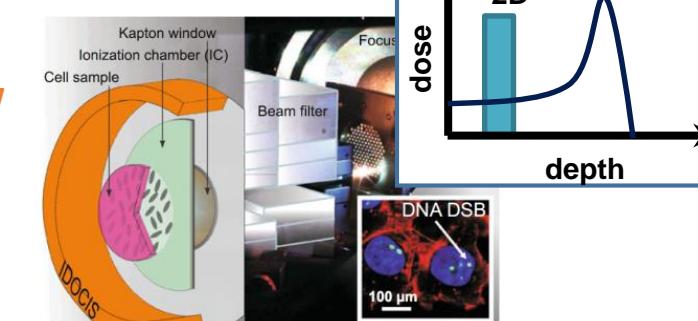


2020 Roadmap on plasma accelerators
New Journal of Physics 23, 031101 (2021)

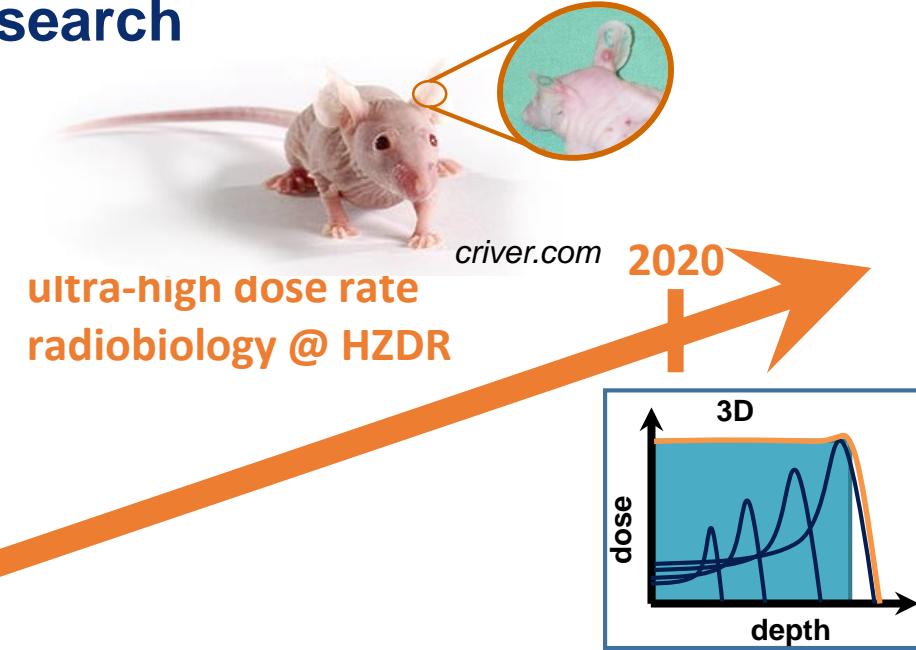
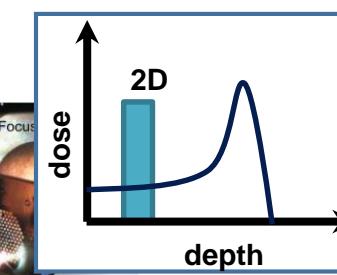
Laser-driven proton acceleration for radiobiological research



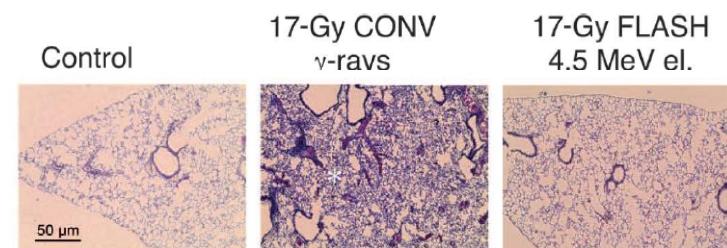
2000
~ 60 MeV protons at
Nova PW (TNSA)
2005 to 2010
20 MeV protons at ~100 TW
Hz rep rate Ti:Sa systems



in vitro radiobiology



Radiation toxicity of RT: **ultrahigh dose-rate FLASH** irradiation increases the differential response between normal and tumor tissue in mice



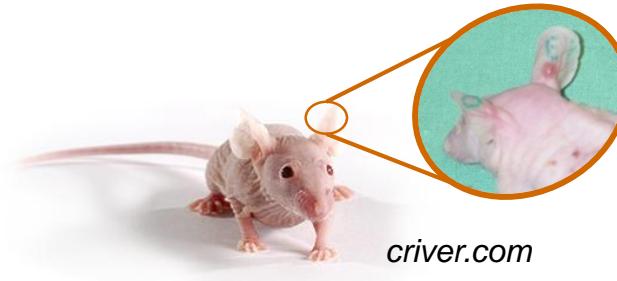
Favaudon et al. Sci. Transl. Med. 2014

compact accelerators for radiotherapy

ultra-high dose rate translational radiobiology

Small animal pilot study with laser-driven proton pulses

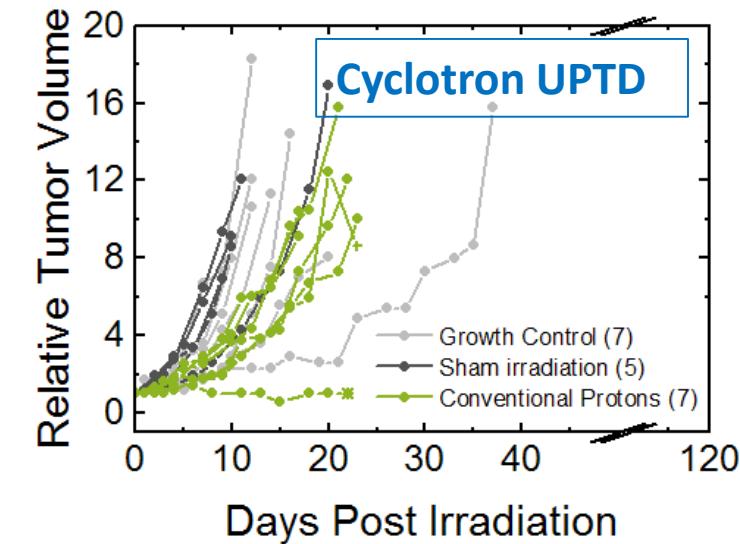
Preparation of comparative *in vivo* radiobiological studies for dose rate effect studies



criver.com

Radiobiological model & requirements:

- radiobiological endpoint: tumor-growth delay of mouse ear tumor
- irradiated volume \varnothing 5 mm, 5 mm depth
- 4 Gy $< +/- 10\%$
- homogeneity $< 10\%$ dev. dose deposition at 4 Gy ($< 10\%$ sample-to-sample variation)
- 2 cohorts (Draco PW & UPTD) with 5 treatment groups each



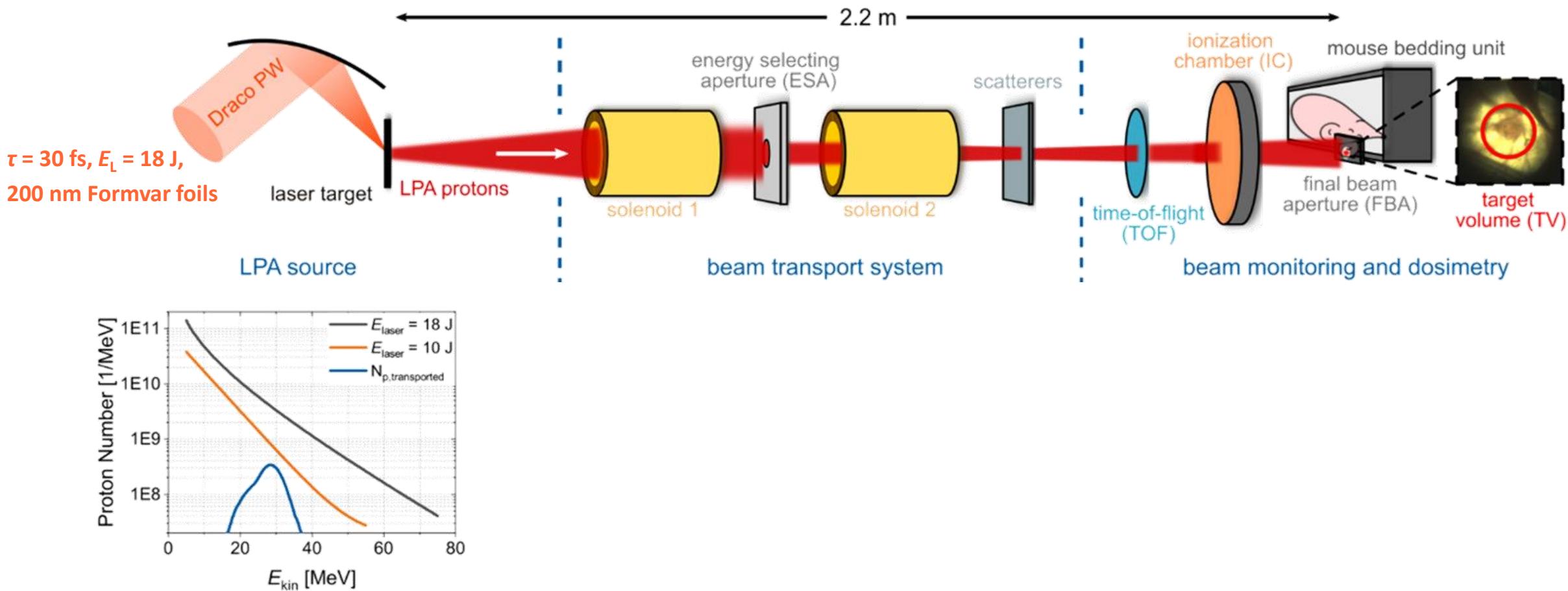
	Draco PW	UPTD
mean dose		3.9
single dose accuracy (2σ)		14%
dose homogeneity lateral/depth (2σ)		9%/2%
mean dose rate		3.6 Gy/min
peak dose rate		-

K. Brüchner et al., Radiat. Onc., Vol. 9 (2014)
Animal study approval DD24-5131/338/35



Small animal pilot study with laser-driven proton pulses

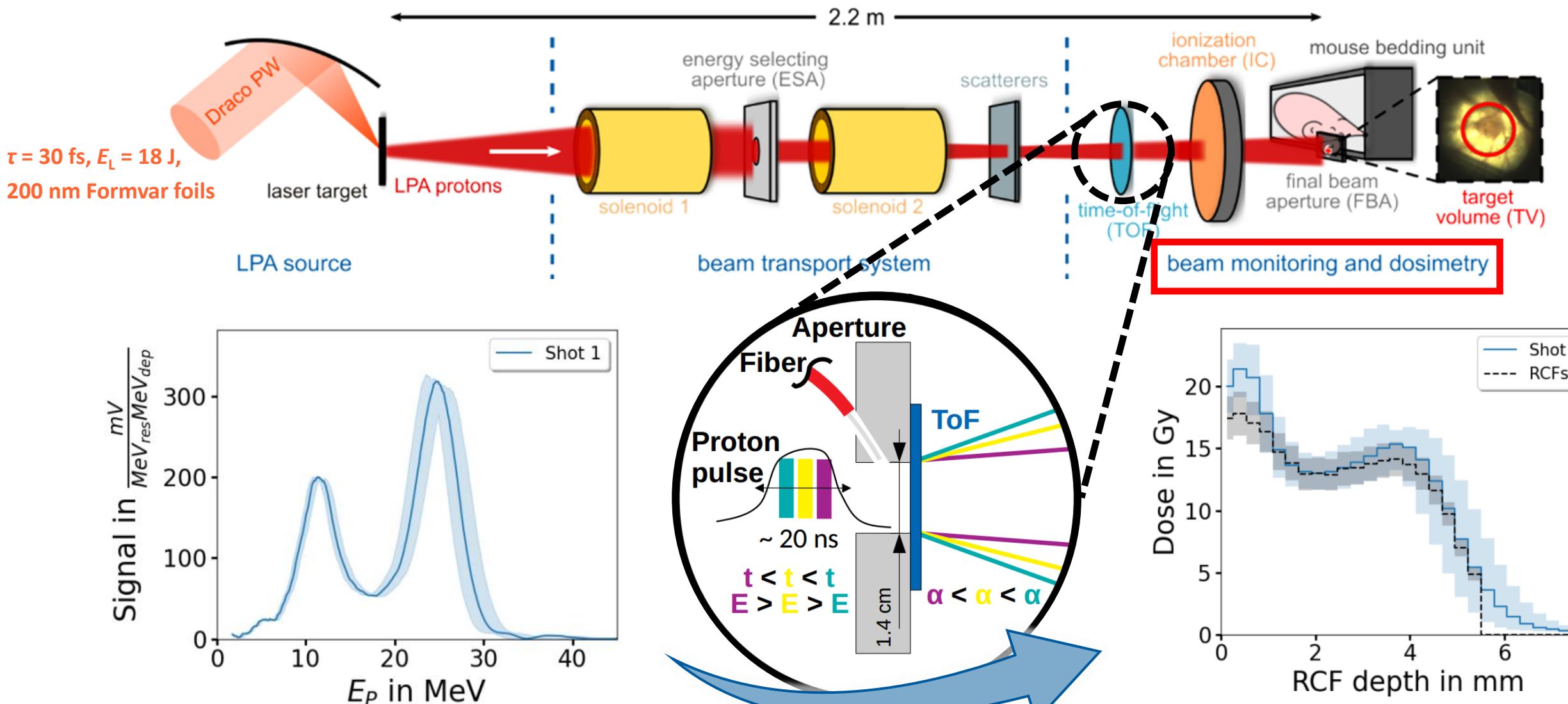
Setup at Draco PW



T. Ziegler *et al.*, Sci Rep 11 (2021); F.-E. Brack *et al.*, Sci Rep 10 (2020), F. Kroll *et al.* Nature Physics (2022)

Small animal pilot study with laser-driven proton pulses

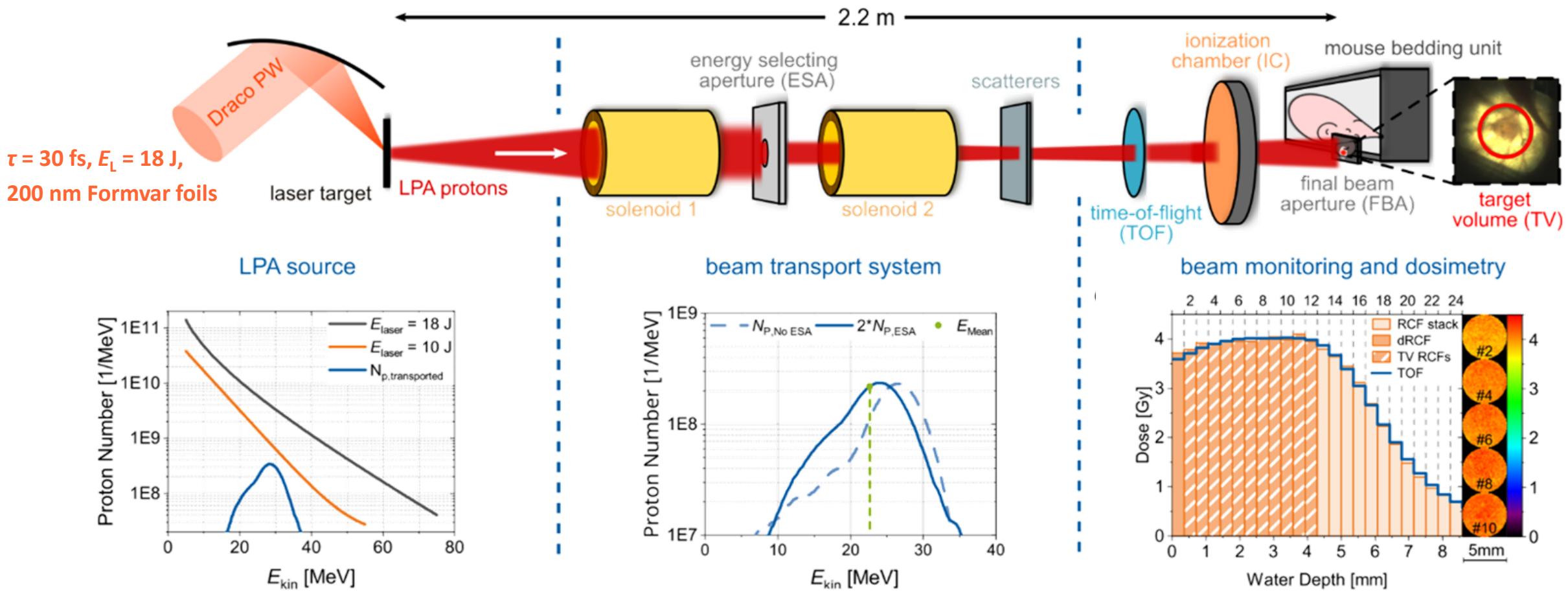
Setup at Draco PW



T. Ziegler *et al.*, Sci Rep 11 (2021); F.-E. Brack *et al.*, Sci Rep 10 (2020), F. Kroll *et al.* Nature Physics (2022), M. Reimold *et al.* in review

Small animal pilot study with laser-driven proton pulses

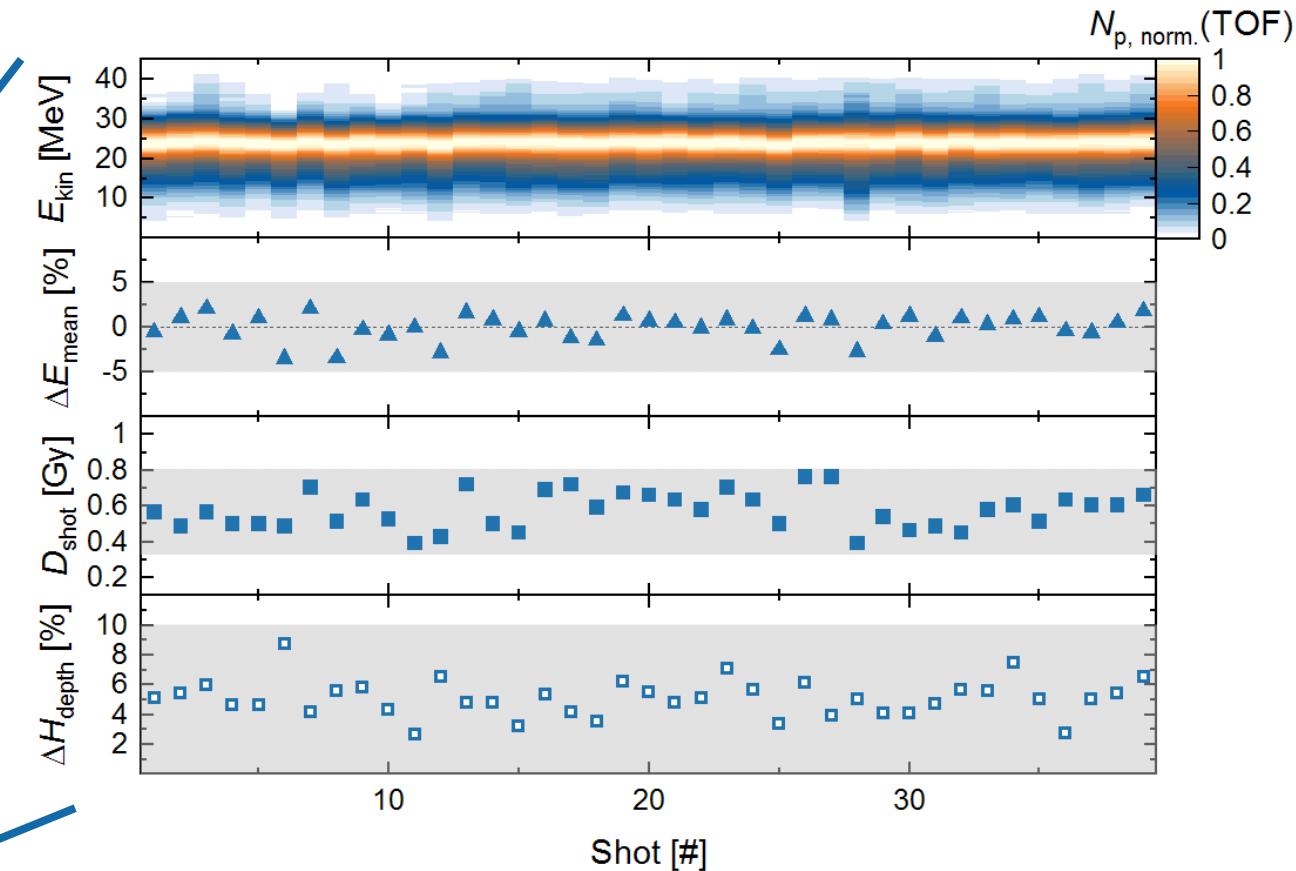
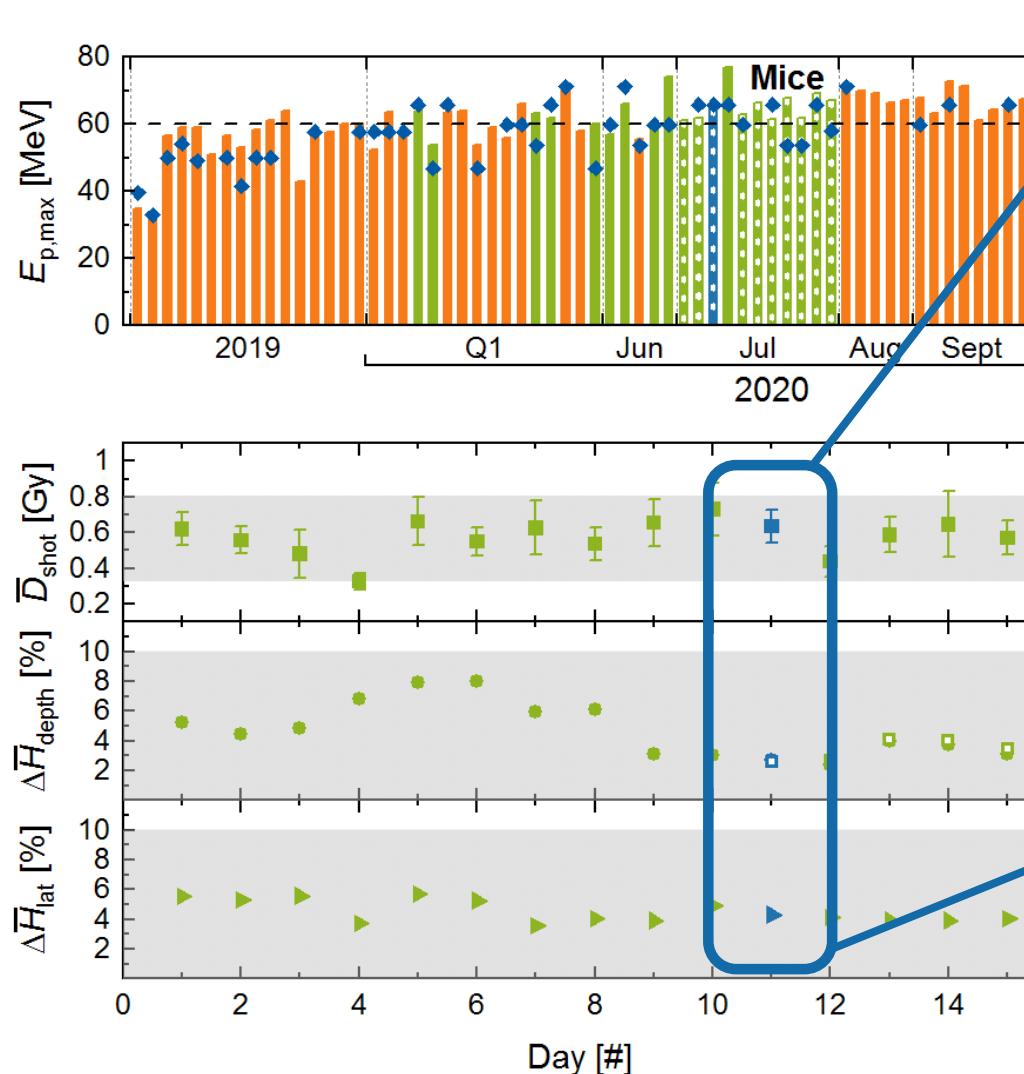
Setup at Draco PW



platform enables single-shot delivery of mm-scale 3D tumor-conform dose distributions making perfect use of the broadband LPA proton spectrum

Small animal pilot study with laser-driven proton pulses

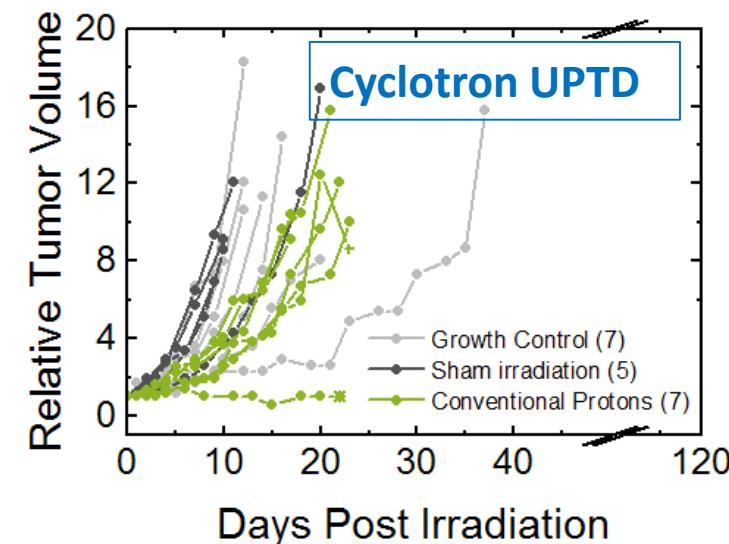
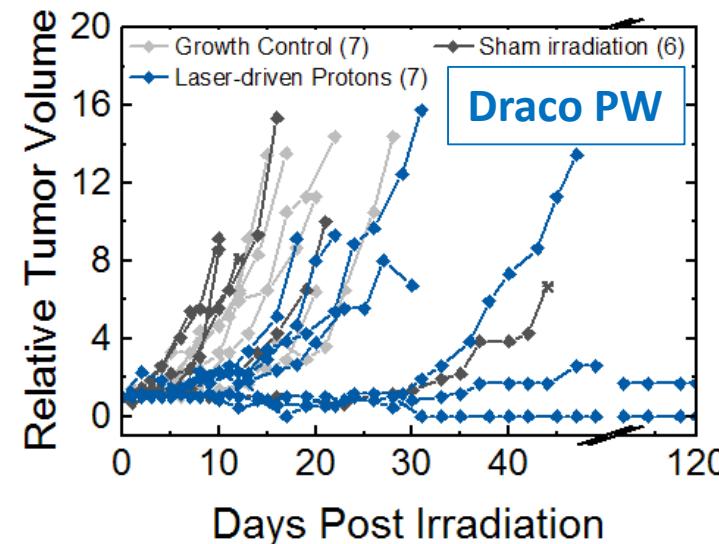
Accelerator readiness and stability benchmarked via application-specific parameters



Small animal pilot study with laser-driven proton pulses

Preparation of comparative *in vivo* radiobiological studies for dose rate effect studies

- model-conform dose delivery
 - ✓ ... mitigation of LPA-inherent spectral intensity fluctuations
- accelerator readiness and stability
 - ✓ ... stable daily accelerator performance over weeks enabling a bio-driven schedule
- radiobiological pilot study
 - ✓ ... meaningful dose-effect data via
 - ✓ ... on-demand proton LPA source operation
 - ✓ ... precise dose delivery & dosimetry
 - ✓ ... complex *in vivo* sample preparation, irradiation & follow-up
- Interesting radiation induced (4 Gy) effect observed, but no significant conclusion because of too small sample number



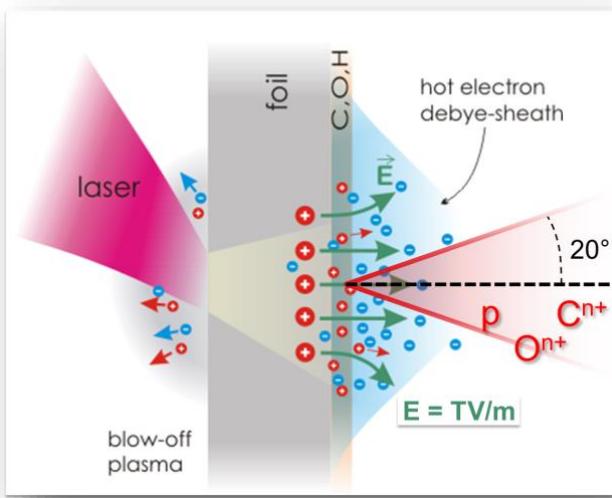
	Draco PW	UPTD
mean dose	3.9	3.9
single dose accuracy (2σ)	8%	14%
dose homogeneity lateral/depth (2σ)	9%/ $< 9\%$	9%/2%
mean dose rate	1.2 – 2.2 Gy/min	3.6 Gy/min
peak dose rate	10^8 Gy/s	-

F. Kroll, et al., Nature Physics 18, 316 (2022)

Upscaling the energy: Enhanced acceleration with near- critical density targets

Tailoring the target (plasma) density profile as decisive parameter

TNSA



mirror-like behaviour
pulse mostly reflected



Target
density

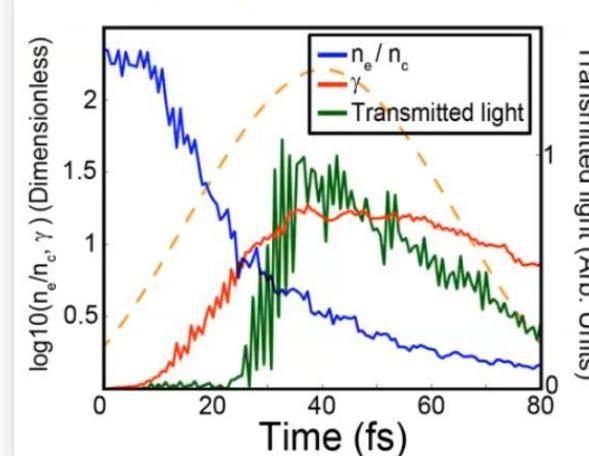
typ. $> 10^{23} \text{ cm}^{-3}$

Dense, opaque target

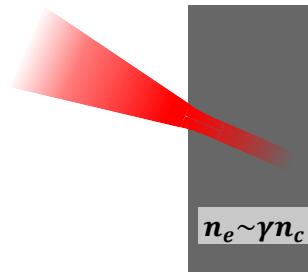
Relativistic transparency

Yin POP (2011), D'humieres POP (2015), Higginson Nat. Comm. (2018),
McKenna, Gonzales-Izquierdo et al. SPIE (2021) & ApplSci (2018)

EPOCH simulation of RIT



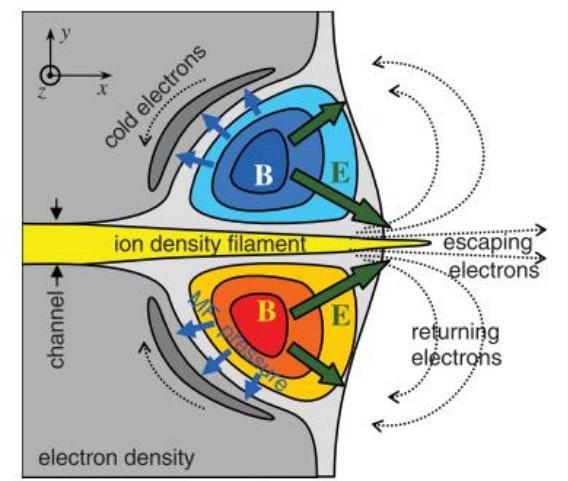
pulse mostly absorbed



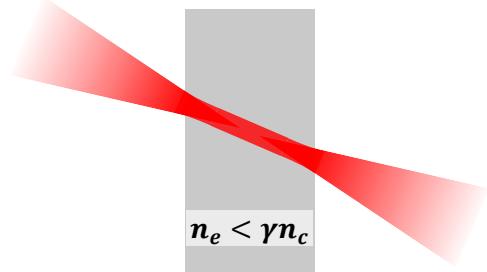
$1.7 \times 10^{21} \text{ cm}^{-3} = n_{\text{critical}}[800\text{nm}]$
near-critical density

Magneto vortex acceleration

Bulanov & Esirkepov PRL (2007)



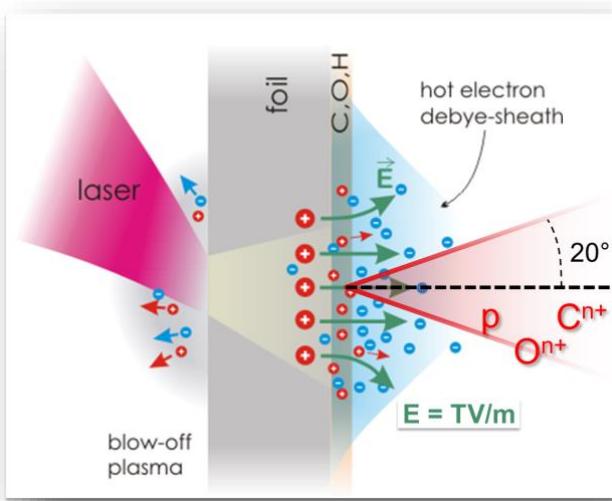
pulse mostly transmitted



typ. $< 10^{20} \text{ cm}^{-3}$
Transparent target

Tailoring the target (plasma) density profile as decisive parameter

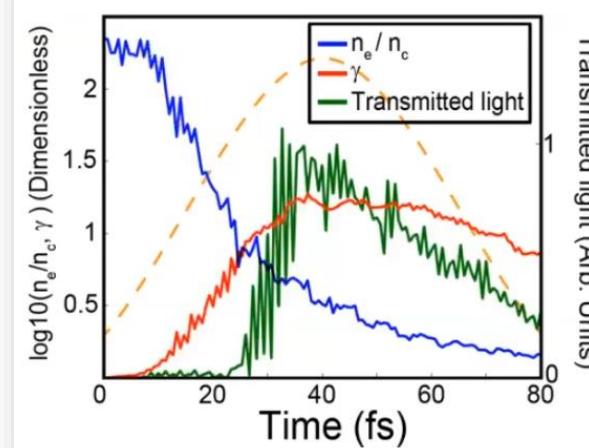
TNSA



Relativistic transparency

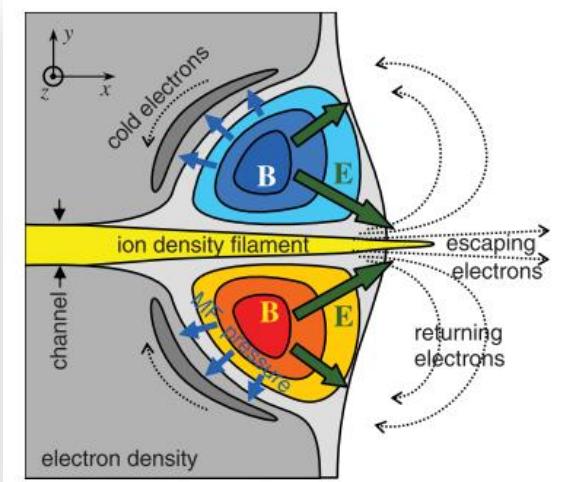
Yin POP (2011), D'humieres POP (2015), Higginson Nat. Comm. (2018),
McKenna, Gonzales-Izquierdo et al. SPIE (2021) & ApplSci (2018)

EPOCH simulation of RIT



Magneto vortex acceleration

Bulanov & Esirkepov PRL (2007)



Solid H_2

Target
density

typ. $> 10^{23} \text{ cm}^{-3}$

Dense, opaque target

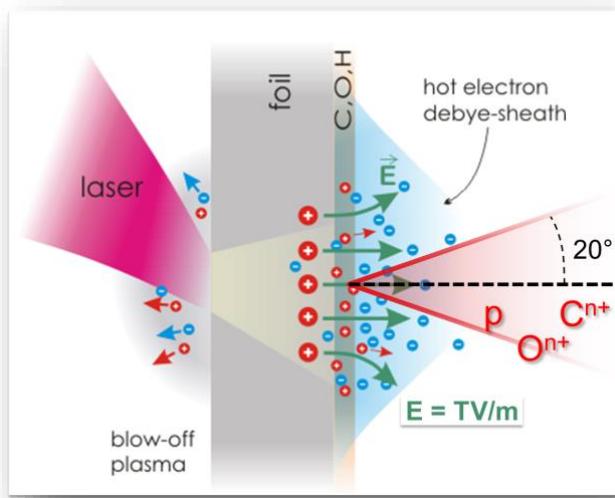
$1.7 \times 10^{21} \text{ cm}^{-3} = n_{\text{critical}}[800\text{nm}]$
near-critical density

typ. $< 10^{20} \text{ cm}^{-3}$

Transparent target

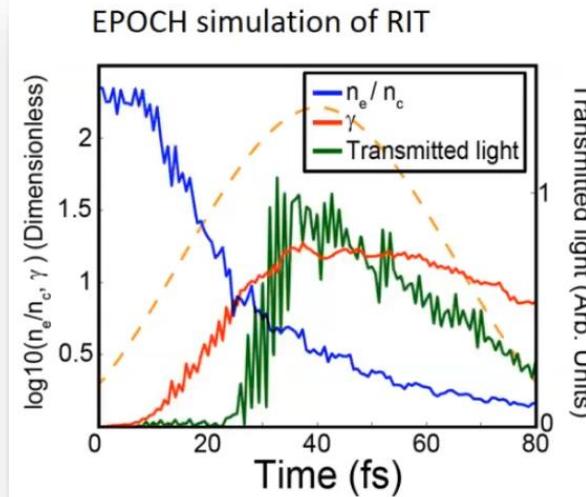
Tailoring the target (plasma) density profile as decisive parameter

TNSA



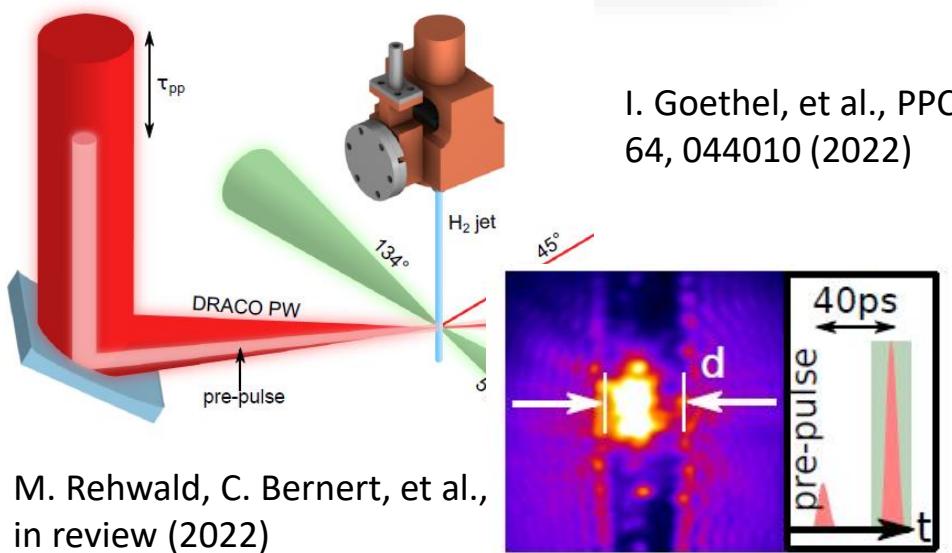
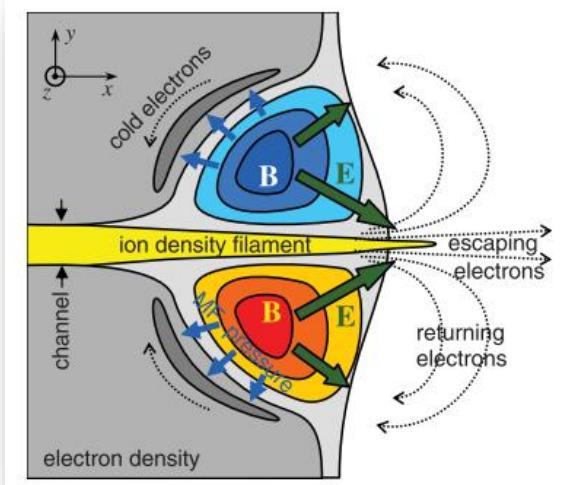
Yin POP (2011), D'humieres POP (2015), Higginson Nat. Comm. (2018),
McKenna, Gonzales-Izquierdo et al. SPIE (2021) & ApplSci (2018)

Relativistic transparency



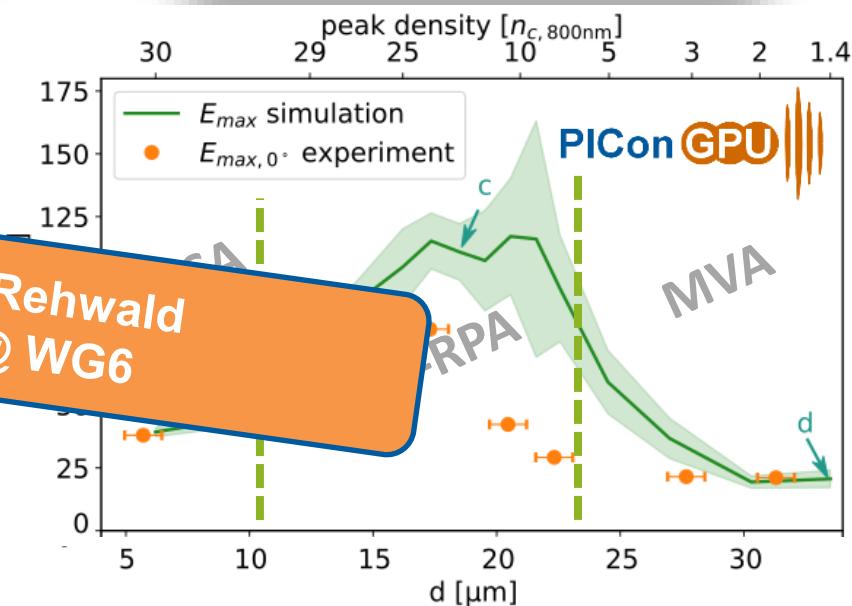
Magneto vortex acceleration

Bulanov & Esirkepov PRL (2007)



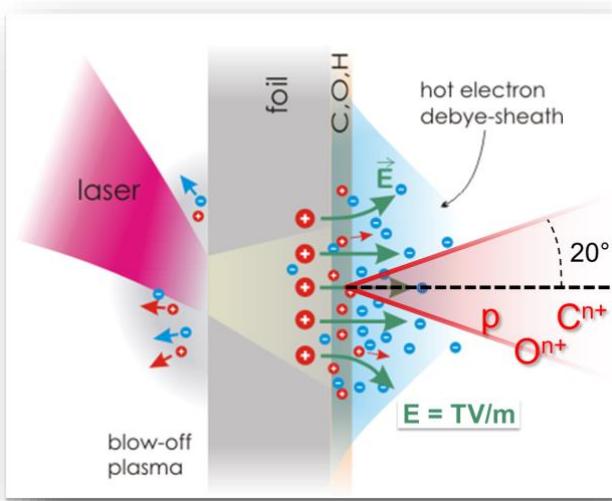
- On-shot hydrogen target density support with designed target enabling quantitative simulation

*Martin Rehwald
Tue @ WG6*



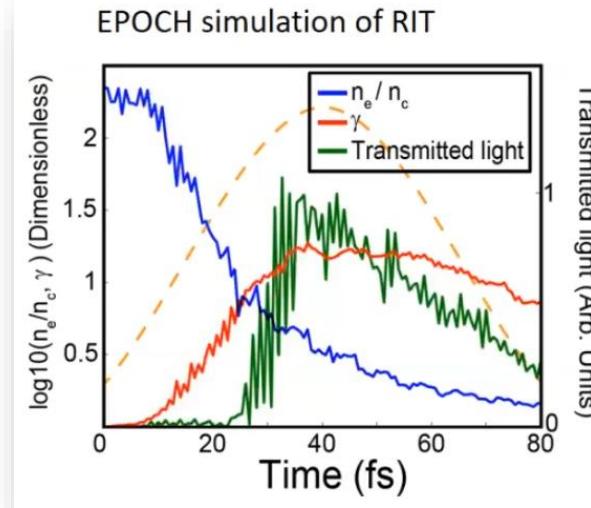
Tailoring the target (plasma) density profile as decisive parameter

TNSA



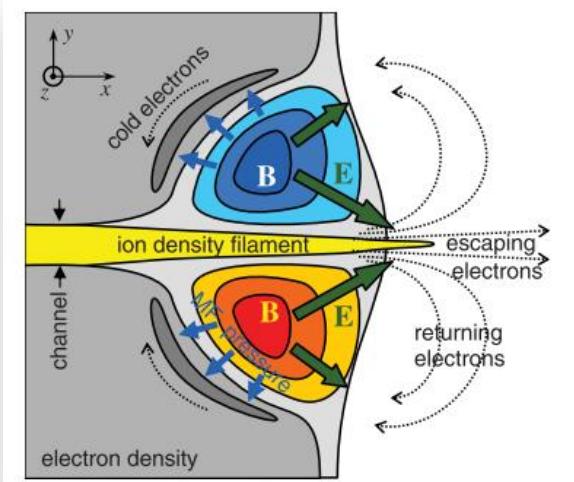
Relativistic transparency

Yin POP (2011), D'humieres POP (2015), Higginson Nat. Comm. (2018),
McKenna, Gonzales-Izquierdo et al. SPIE (2021) & ApplSci (2018)



Magneto vortex acceleration

Bulanov & Esirkepov PRL (2007)



foils

Solid H₂

Target
density

typ. $> 10^{23} \text{ cm}^{-3}$

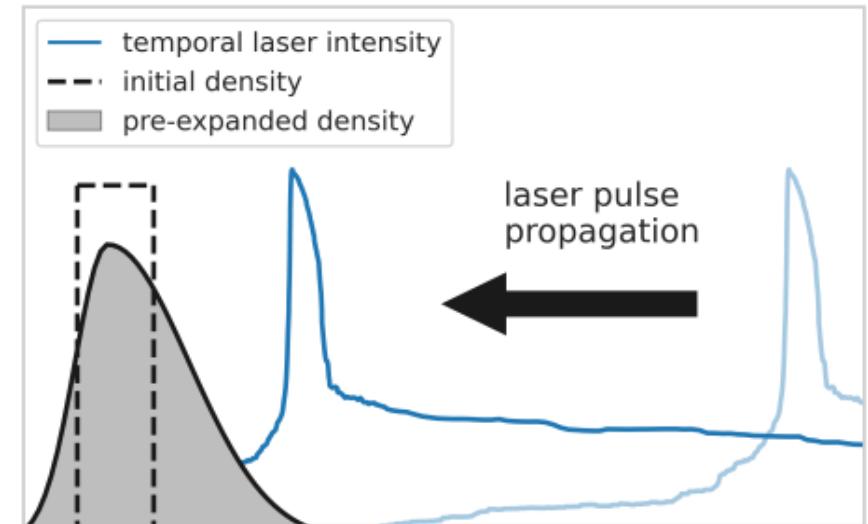
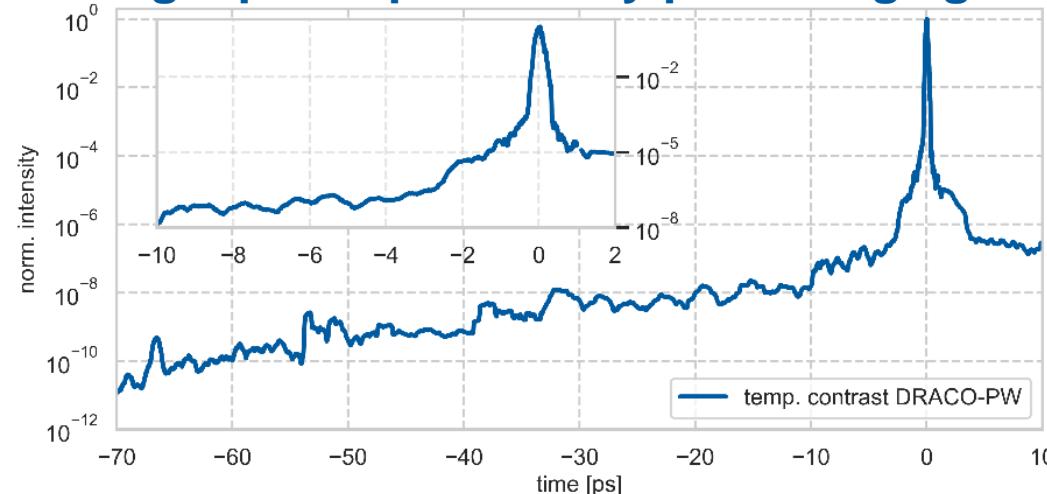
Dense, opaque target

$1.7 \times 10^{21} \text{ cm}^{-3} = n_{\text{critical}}[800\text{nm}]$
near-critical density

typ. $< 10^{20} \text{ cm}^{-3}$

Transparent target

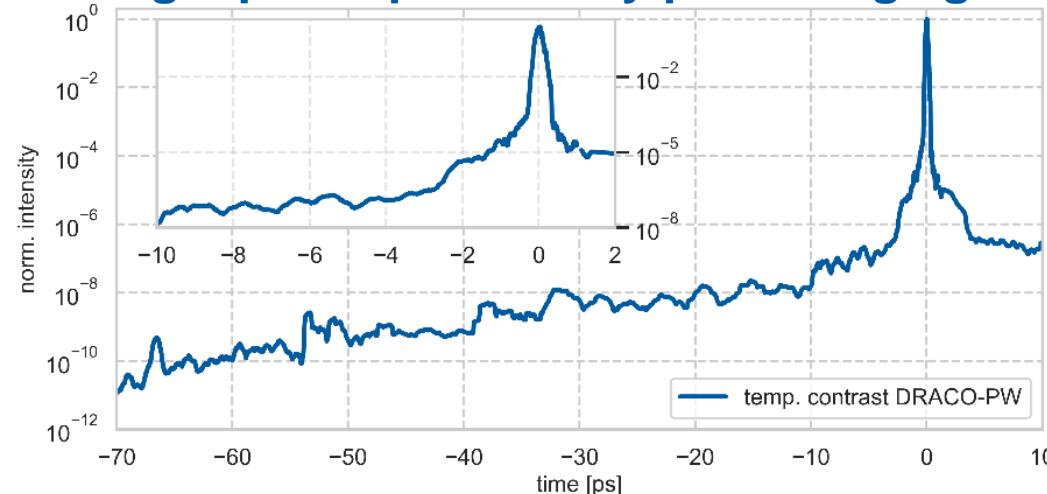
target pre-expansion by preceding light



- targets can get relativistic transparent during interaction
→ increase absorption + coupling into ions + energies

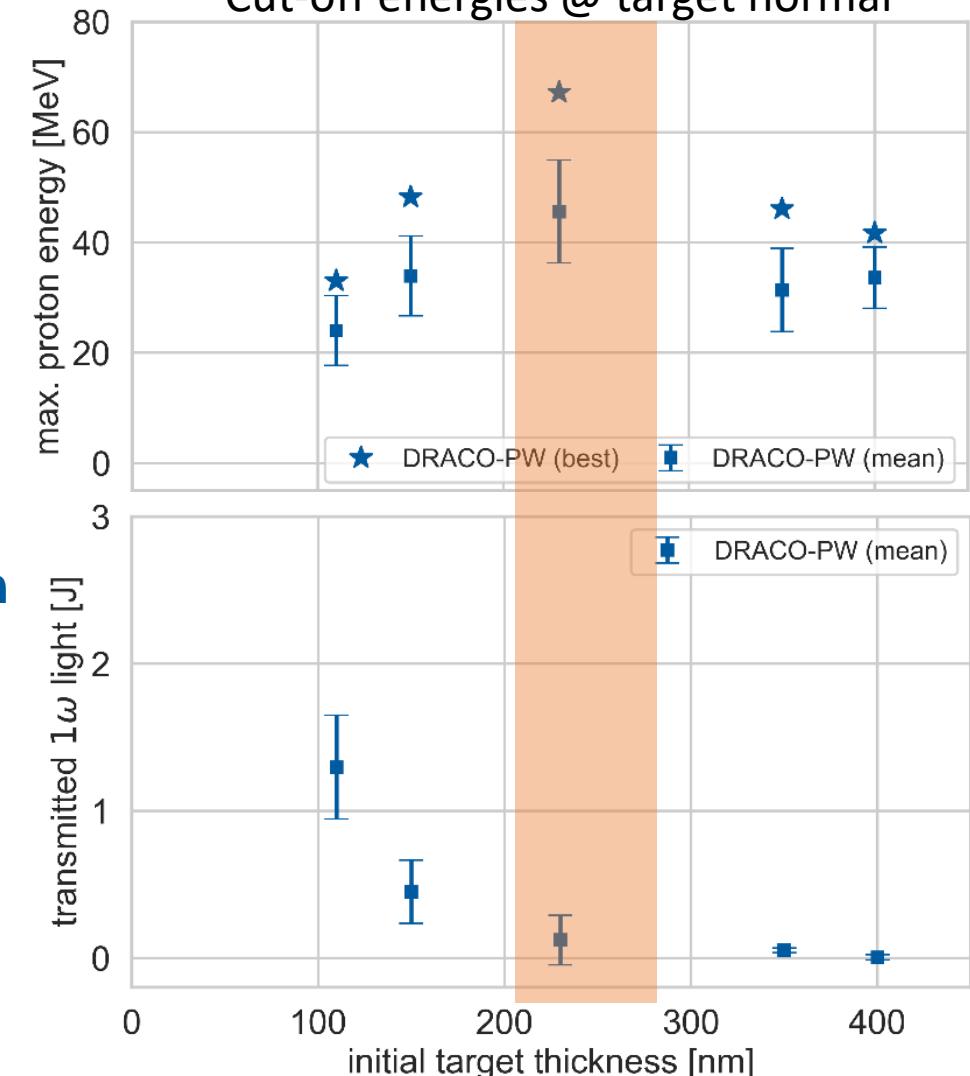
Enhancing ion energies with relativistically transparent targets

target pre-expansion by preceding light



- targets can get relativistically transparent during interaction
→ increase absorption + coupling into ions + energies
- enhanced proton energies at optimal thickness
- onset of transparency at optimal thickness
- observation of forward acceleration

Cut-off energies @ target normal



Experimental setup

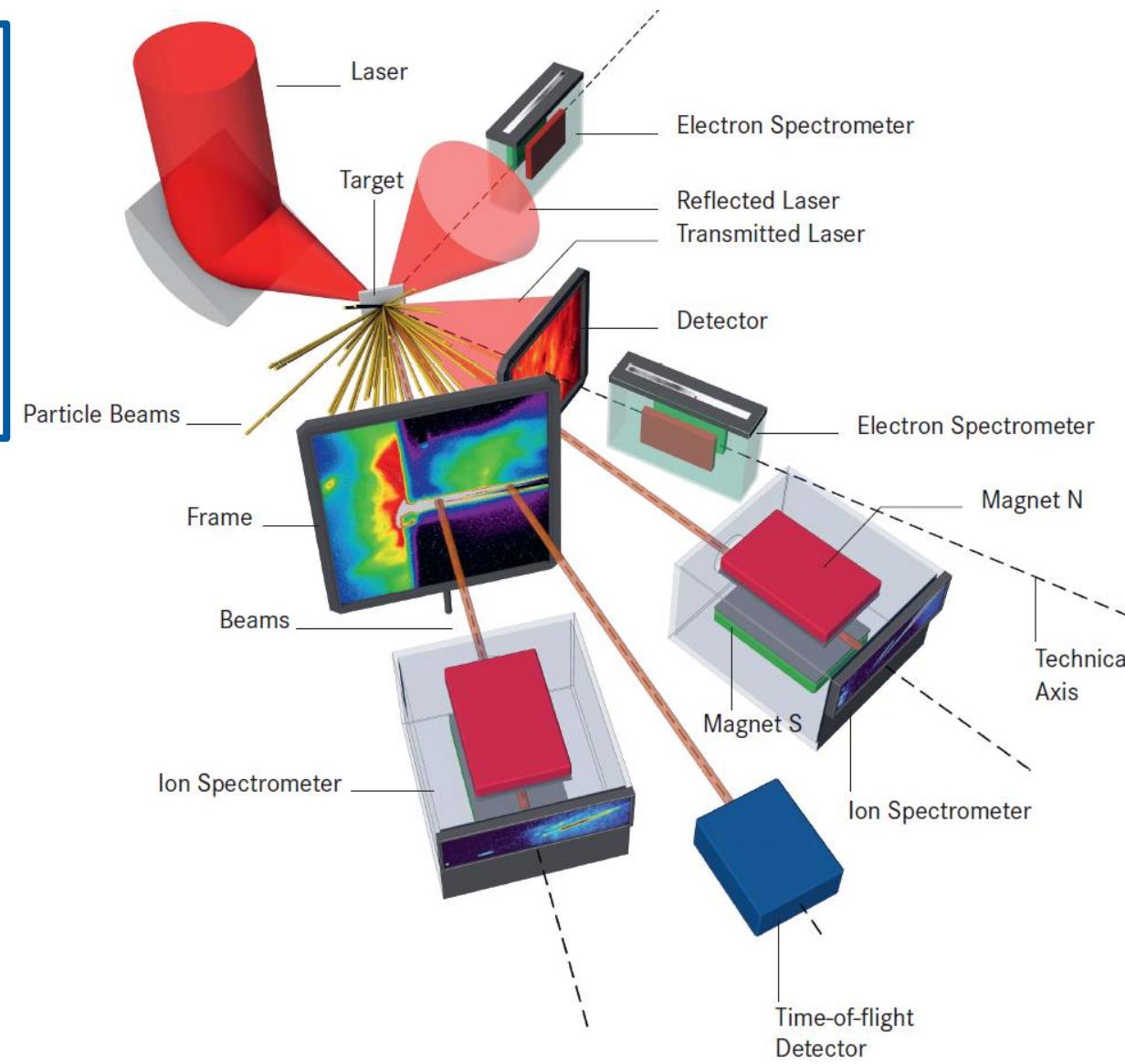
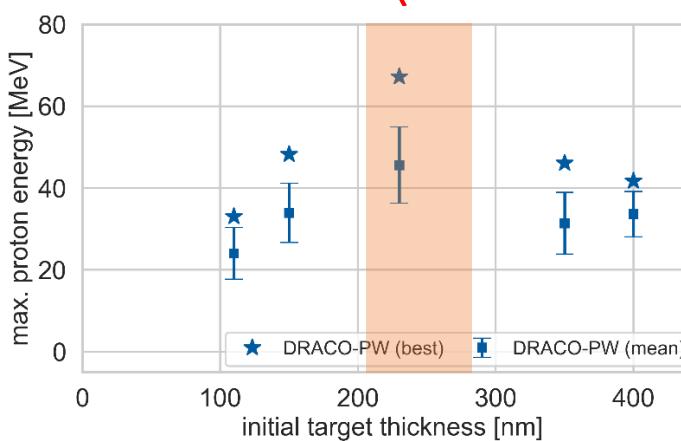
DRACO-PW

$E_L = 22.4 \text{ J (on target), } \tau = 30 \text{ fs}$

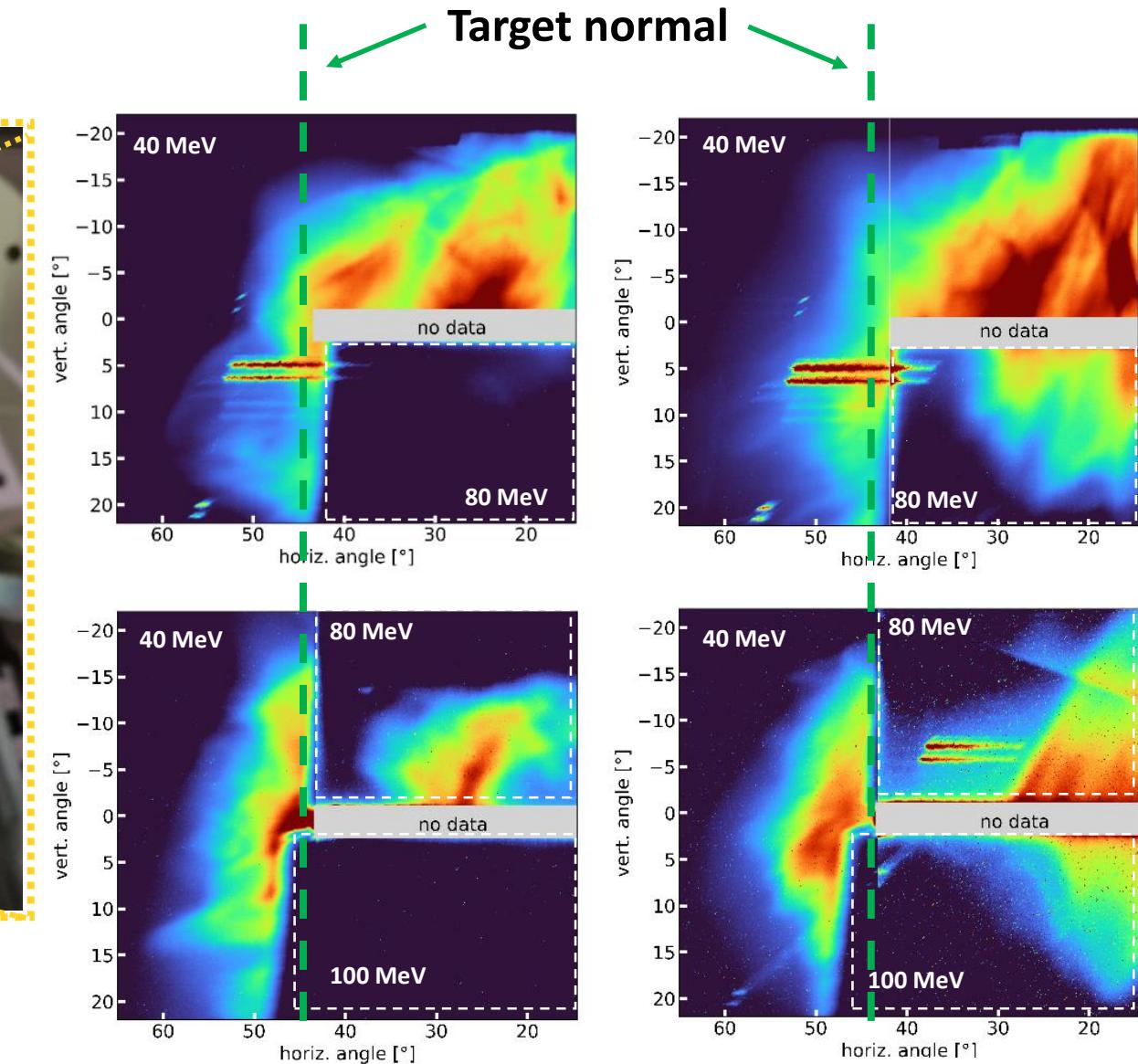
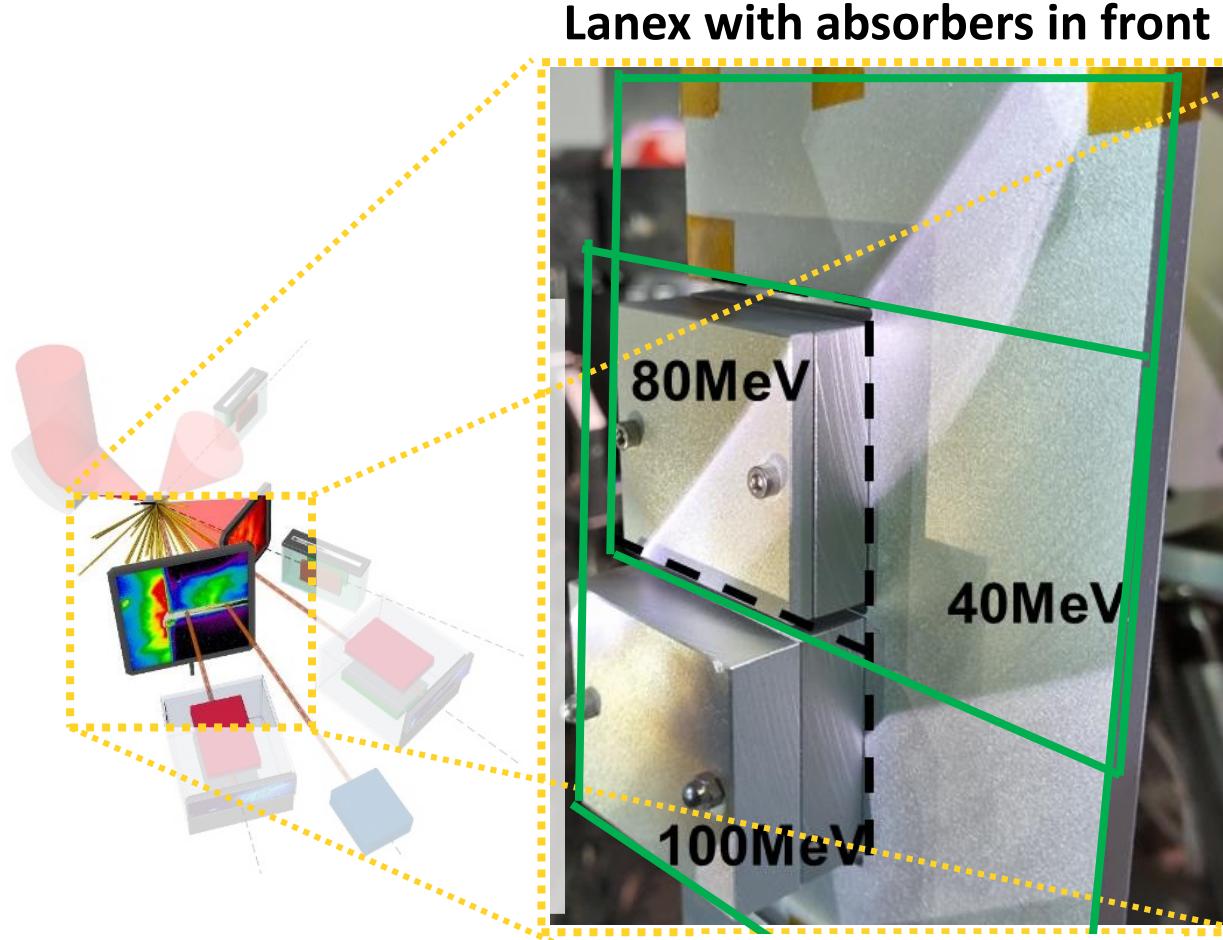
$F/2.3 \text{ OAP} \rightarrow 2.3 \mu\text{m (FWHM)}$

$I_{\text{peak}} = 6.5 \cdot 10^{21} \text{ W/cm}^2 \text{ (a}_0=55)$

targets: $240 \pm 20 \text{ nm Formvar foils}$



Profiler: shift of acceleration direction towards laser axis



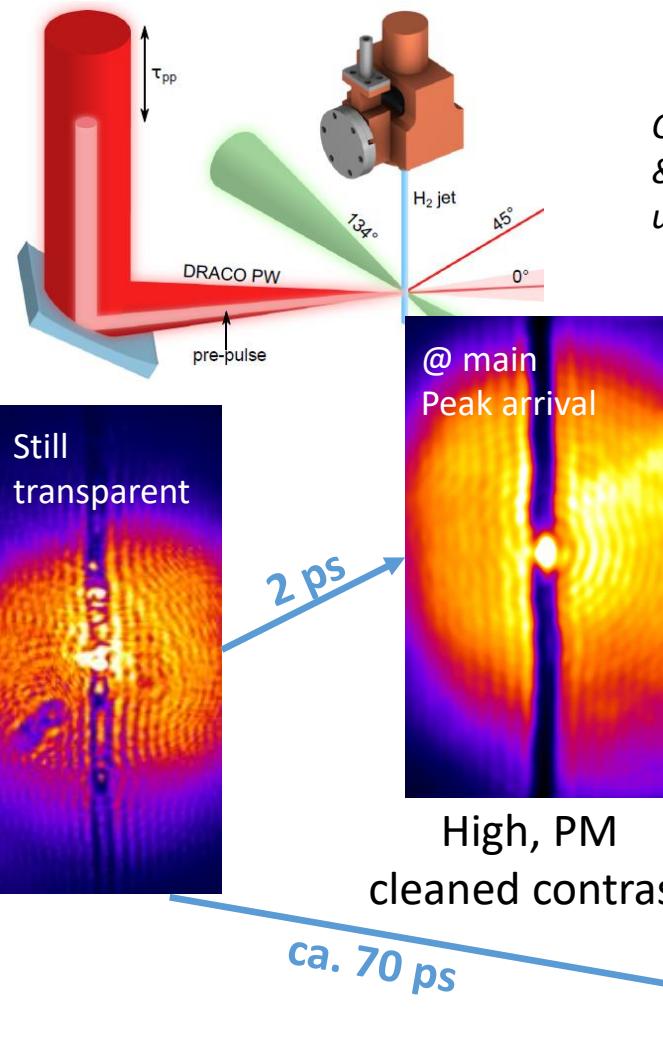
- highest energies in forward direction

T. Ziegler, et al. unpublished

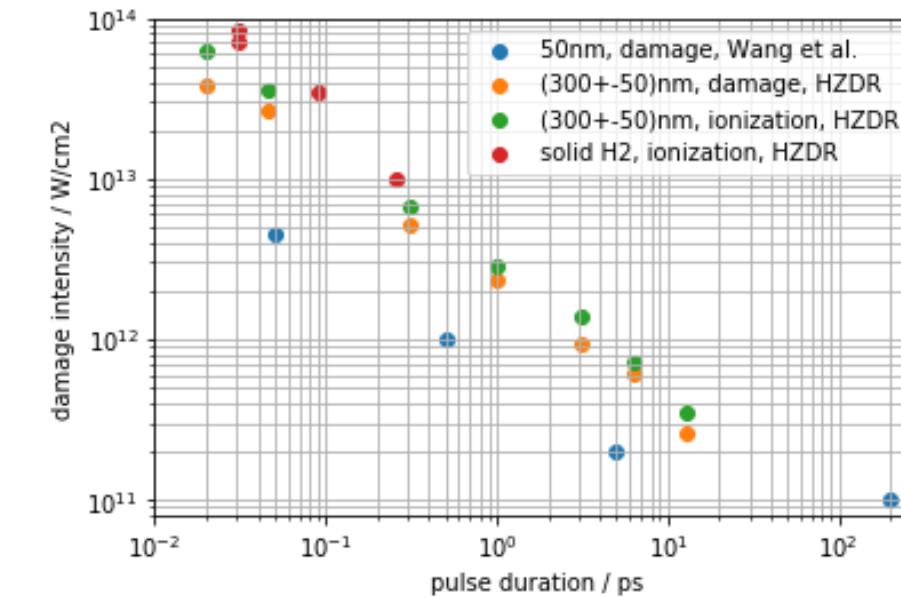
Microscopic understanding and robust performance

Characterization of the laser induced breakdown

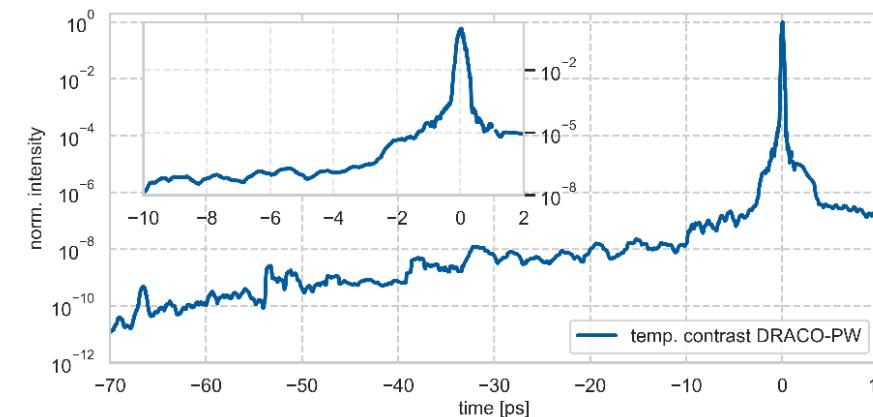
... precisely measure the threshold intensities at low intensities,



C. Bernert et al. SciRep 12, 7278 (2022)
& C. Bernert et al. Phys. Rev. Applied, under review (2022)



... and correlate to the full energy temporal laser contrast (what is real).



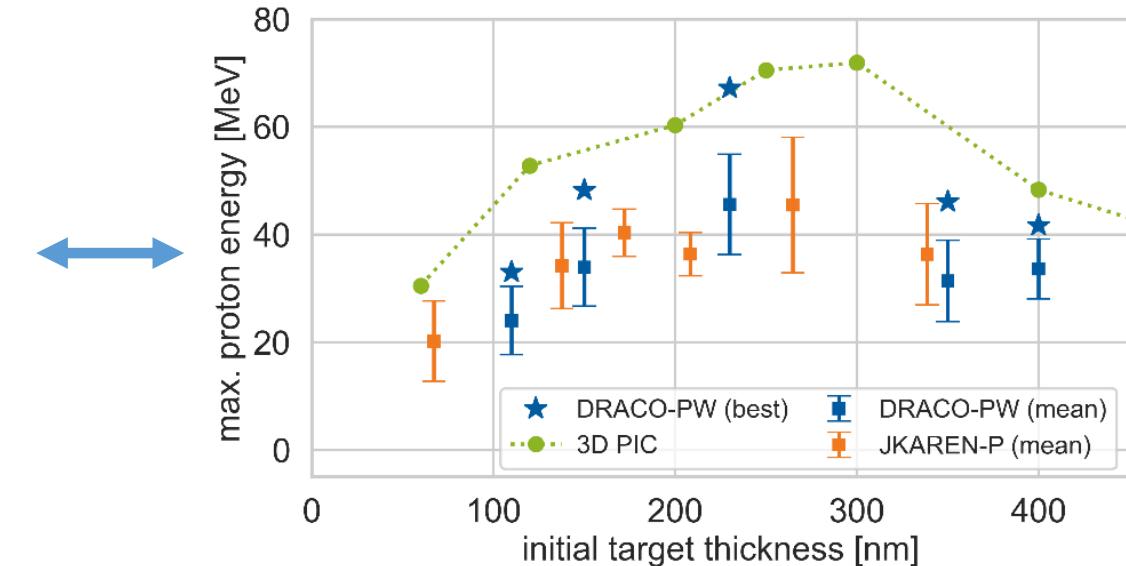
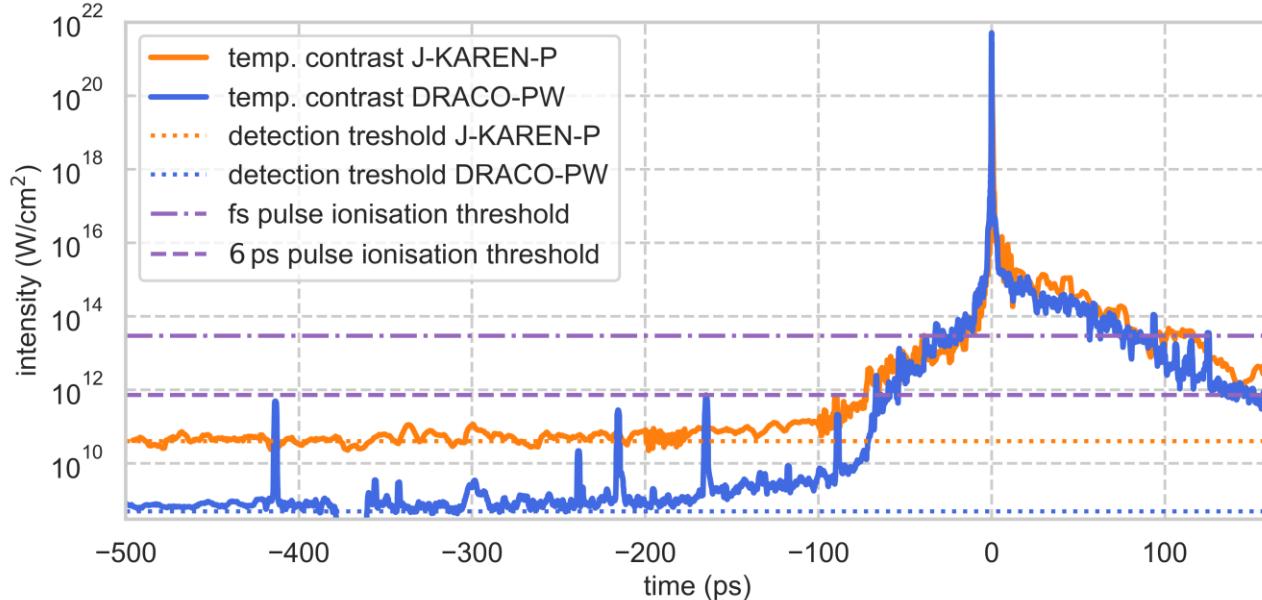
→ Input for start-to-end simulation frameworks based on MHD and 3DPIC.

We can characterize the start of the interaction at full energy laser shots,

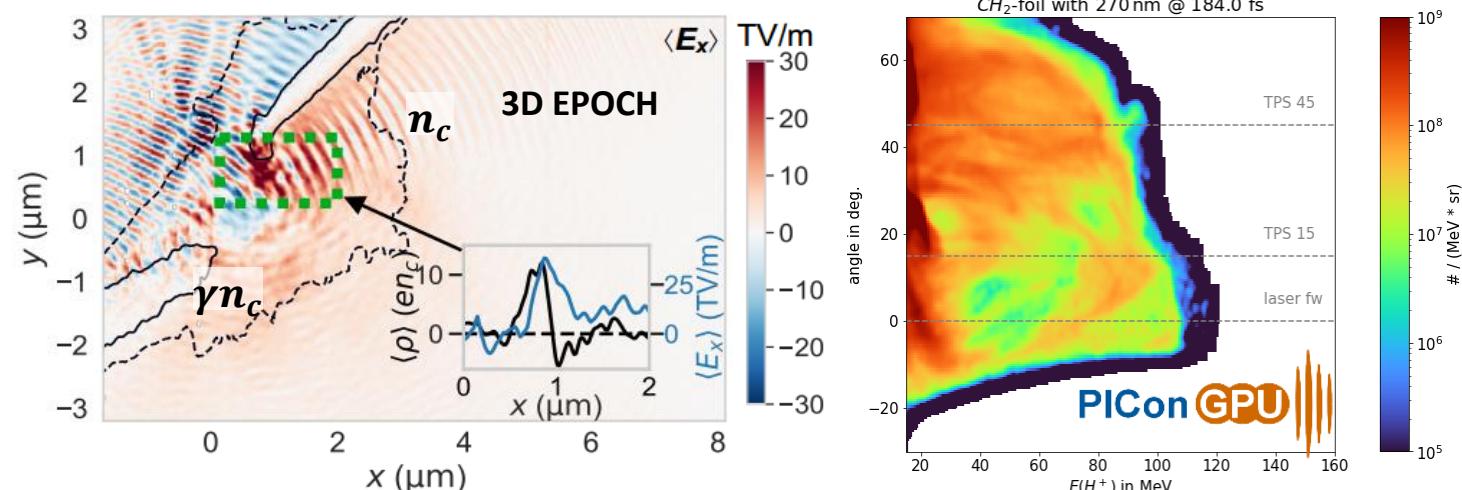
Microscopic understanding and robust performance

Comparing pre-expanded thin foil studies at two laser systems w/o PM

Coordinated laser and plasma diagnostics provide similar results



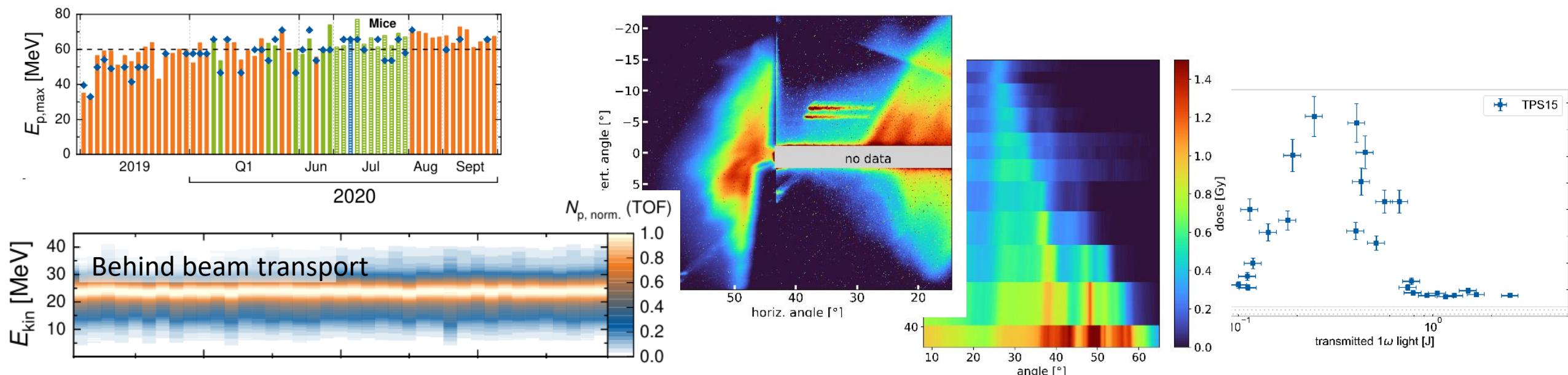
... and enable the establishment of start-to-end simulation framework based on MHD sims and two 3D PIC codes.



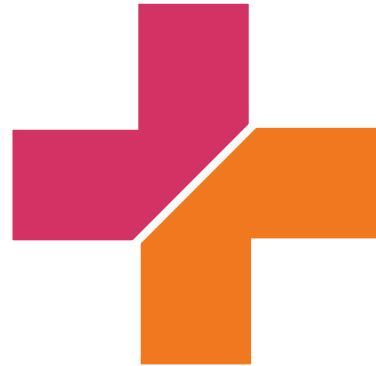
N. Dover, T. Ziegler, et al. in review (2022)

Conclusion

1. Stable TNSA beam performance enabled first volumetric in-vivo irradiation of mice
2. Combining multiple detector systems based on different principles to confirm energies >100 MeV in pre-expanded foils
3. High energy, bandwidth limited pulses from rep-rate capable laser system
4. Understand and mitigate strong fluctuation is work in progress



Big Thanks to the Team and Collaborators



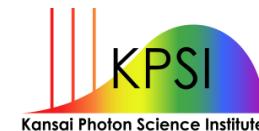
Laser radiooncology

J. Pawelke, E. Beyreuther, K. Brüchner, E. Bodenstein, L. Karsch, E. Lessmann, M. Krause, E. Troost, N. Cordes, C. Richter, et al.
K. Zeil, J. Metzkes-Ng, F. Kroll, S. Assenbaum, C. Bernert, F. Brack, E. Beyreuther, L. Gaus, S. Kraft, A. Nossula, M.E.P. Umland, M. Rehwald, M. Reimold, M. Vescovi, H.-P. Schlenvoigt, M. Sobiella, T. Ziegler, T. Kluge, I. Goethel, S. Bock, R. Gebhardt, U. Helbig, T. Püschen, U. Schramm, T. Cowan, et al.

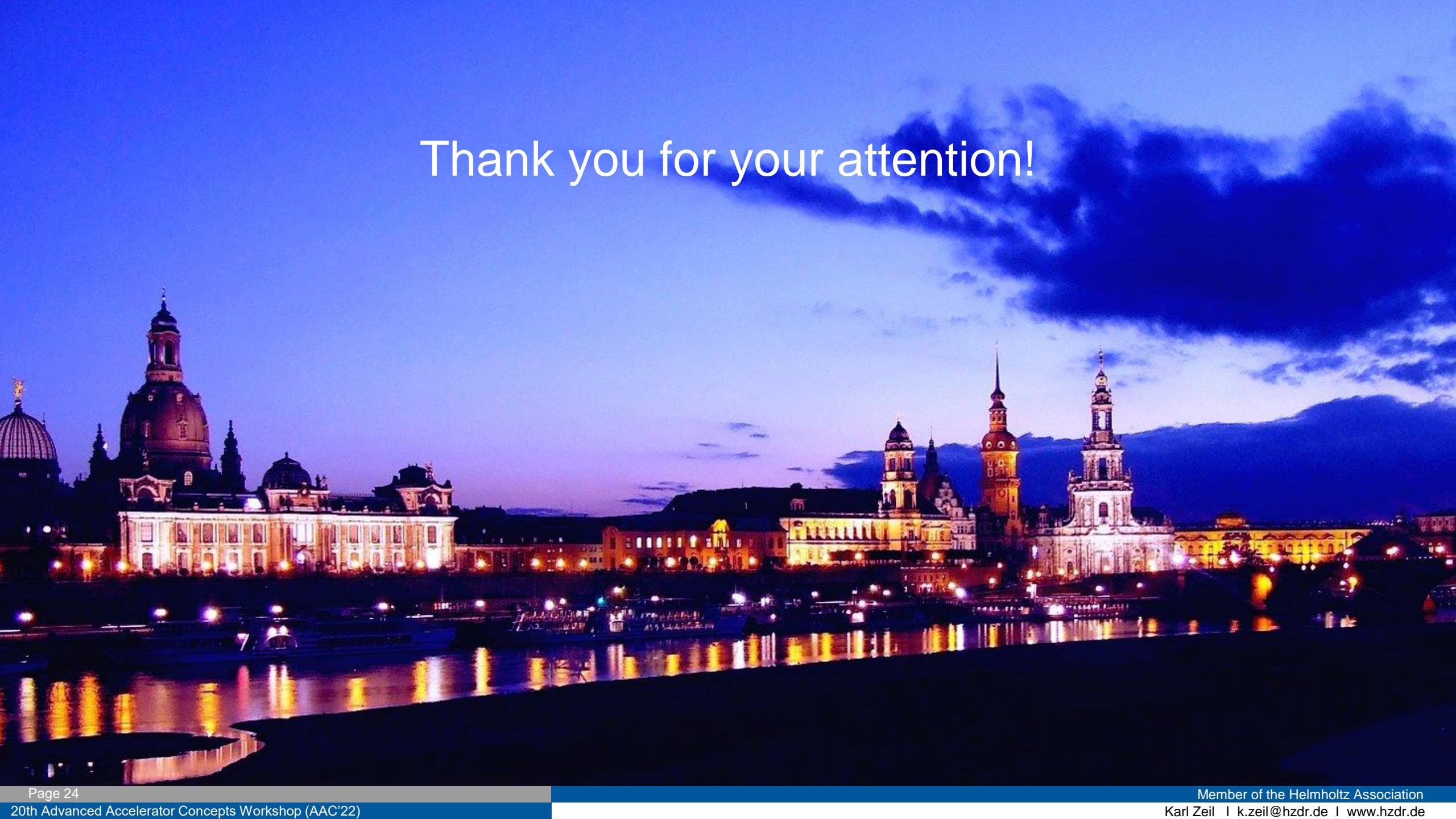
High-field laboratory Dresden (HLD) and HZDR workshop; R. Szabo, et al. (ELI-ALPS); J. Jansen, et al. (DKFZ)



S. Glenzer, C. Curry, M. Gauthier,
J. Kim, F. Fiuza
S. Goede et al.



M. Nichiuchi, H. Kyriama, N. Dover, A. Kon



Thank you for your attention!