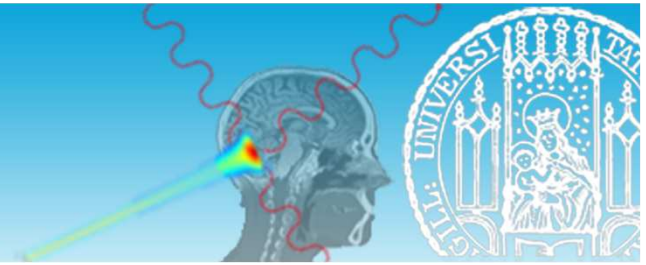




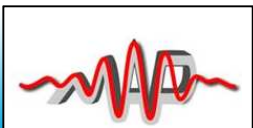
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Detection of laser-accelerated ions

S. Reinhardt

Department of Medical Physics, LMU München

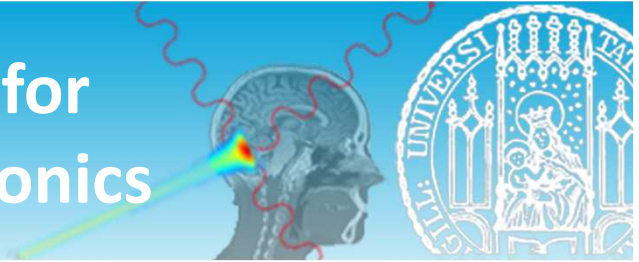




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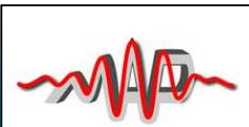
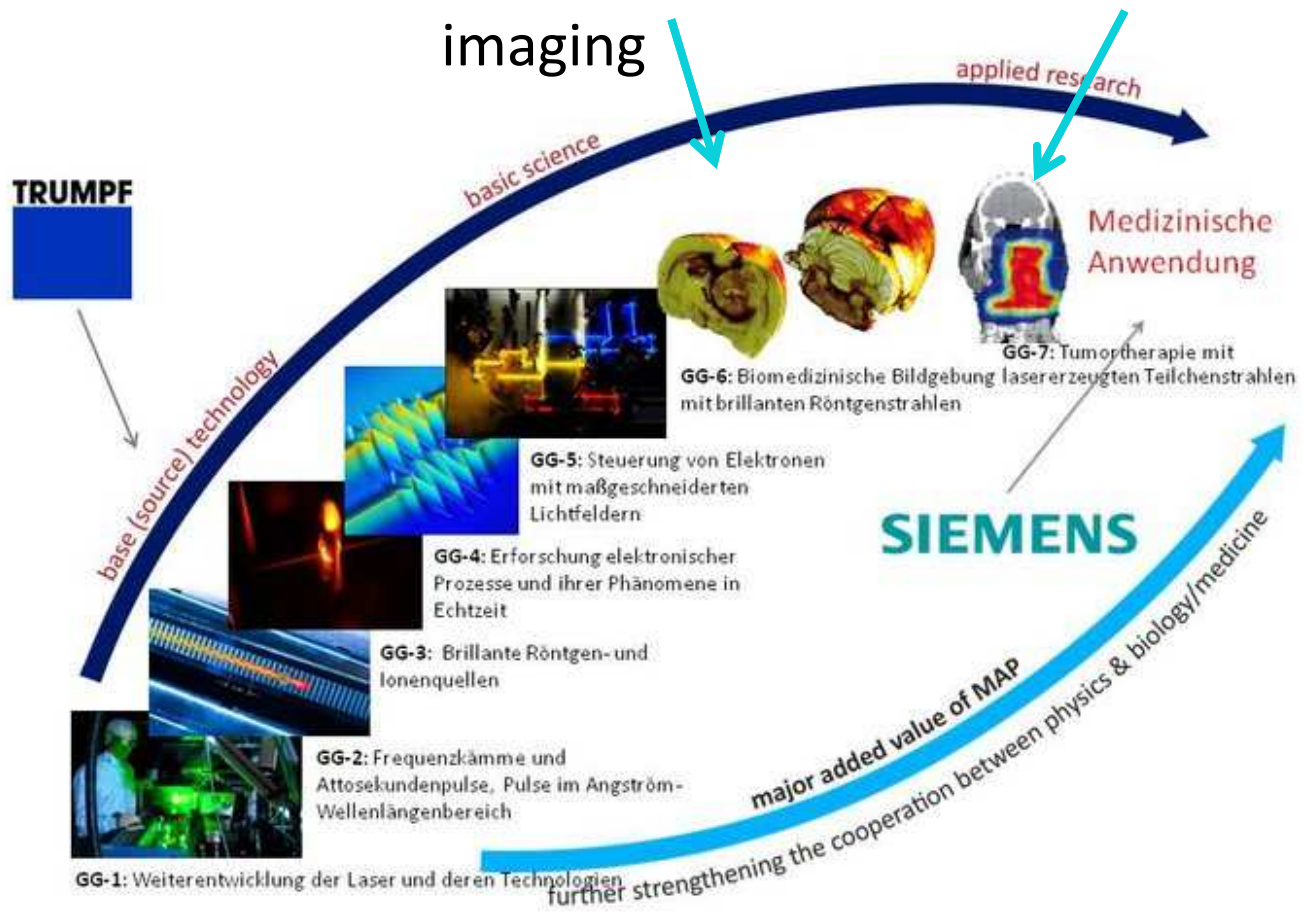
Munich Centre for Advanced Photonics



Medical applications:

Brilliant X-rays for biomedical imaging

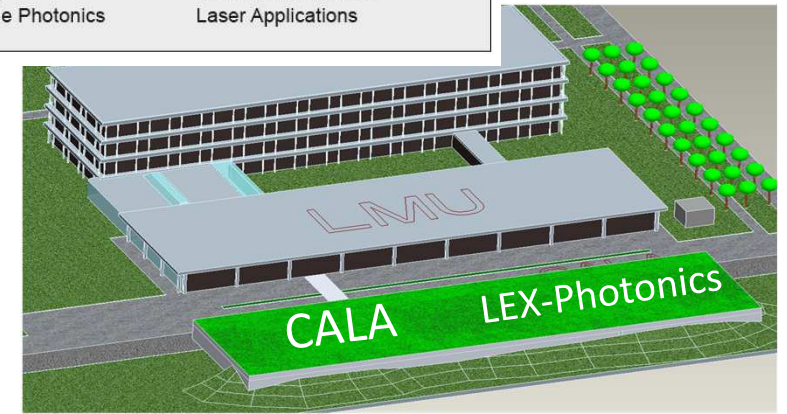
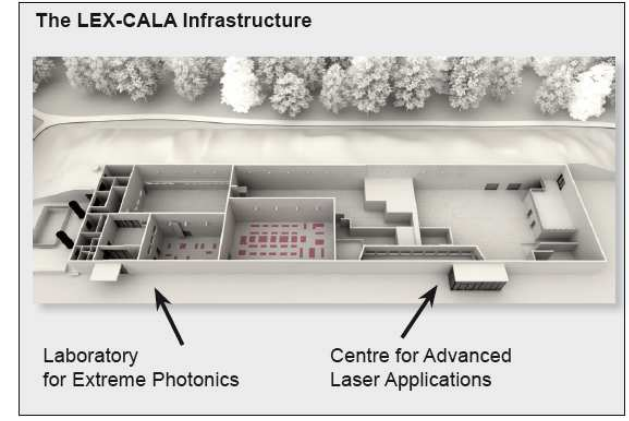
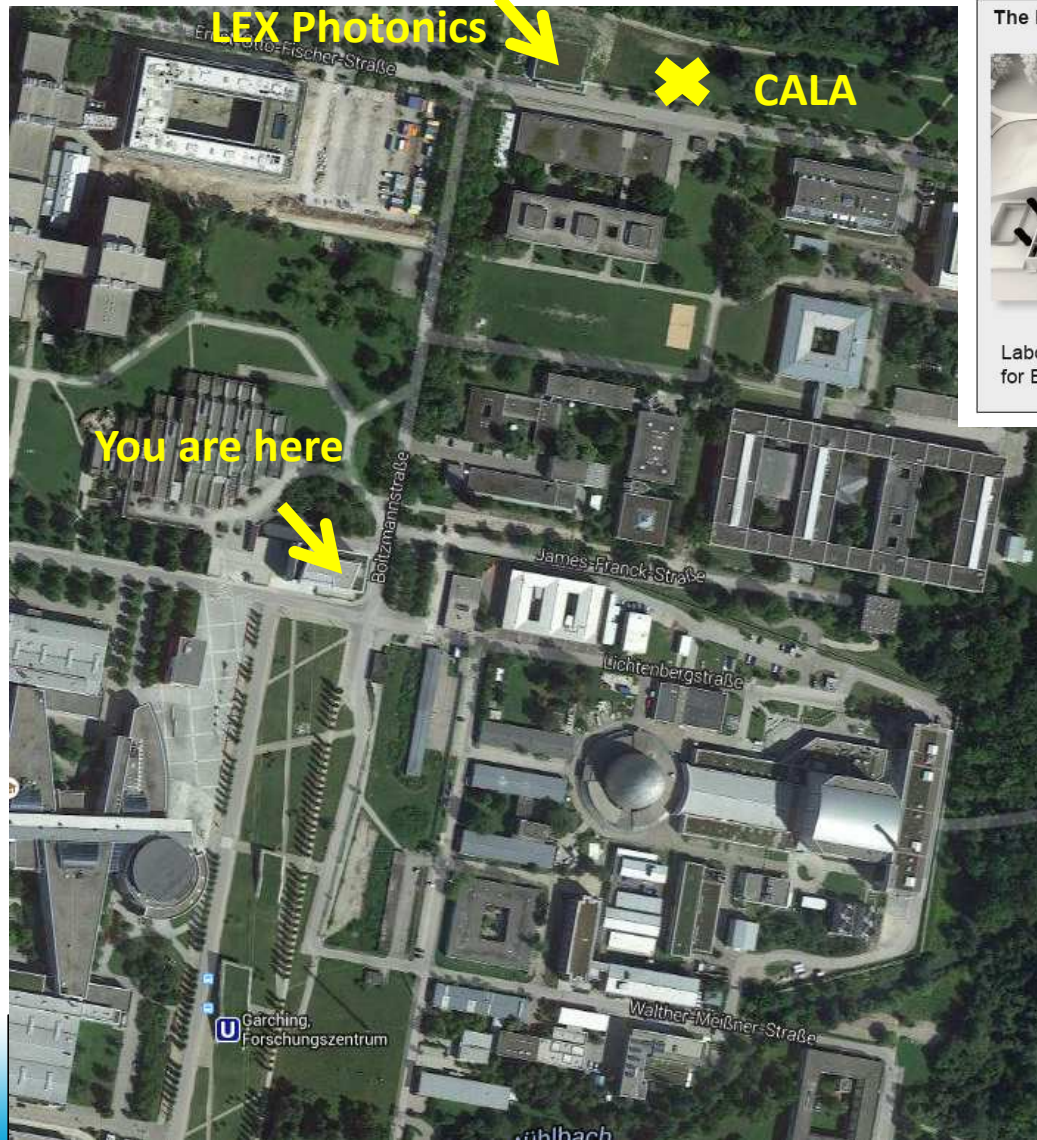
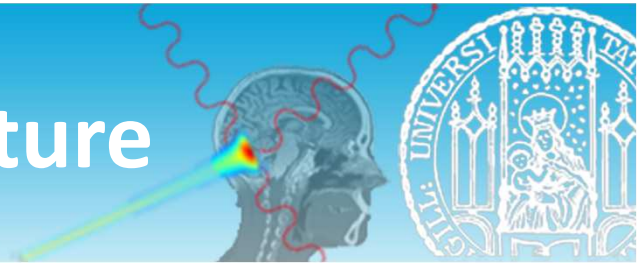
Radiation therapy with laser accelerated particles





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LEX/CALA: infrastructure

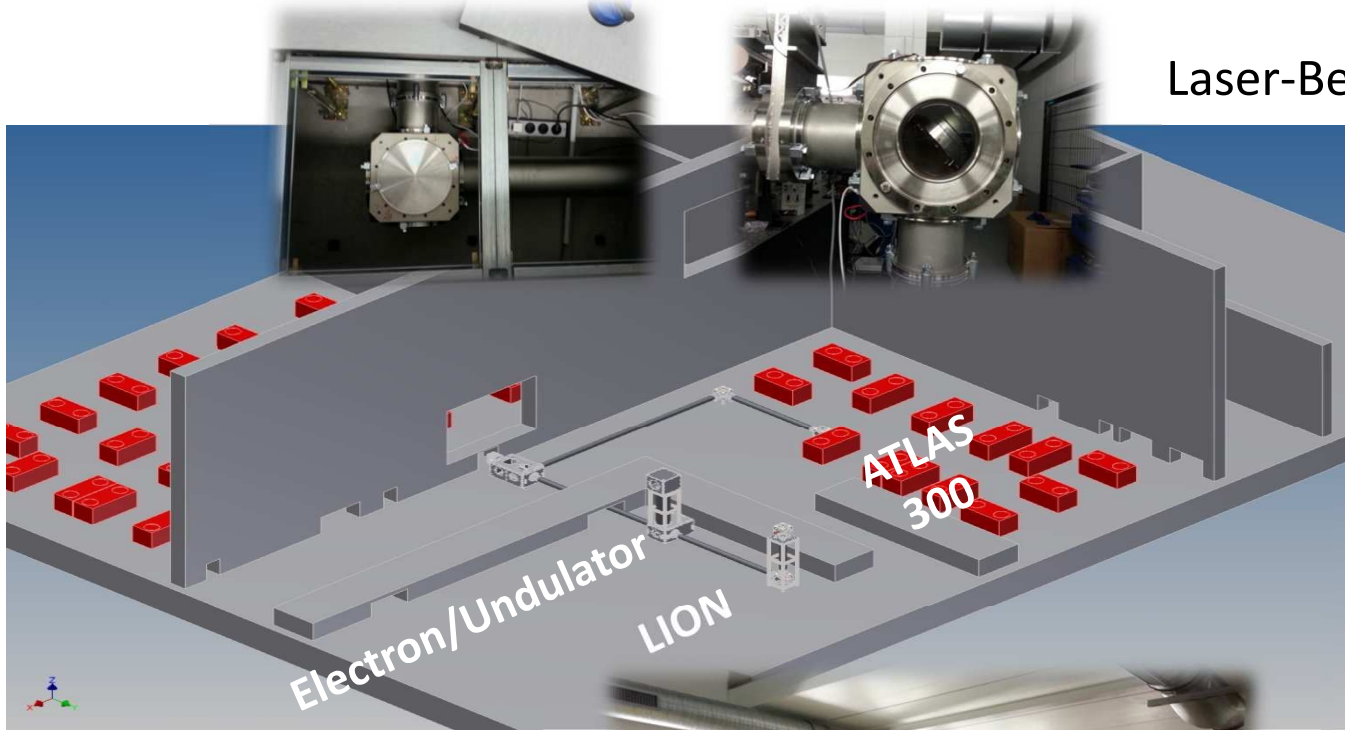
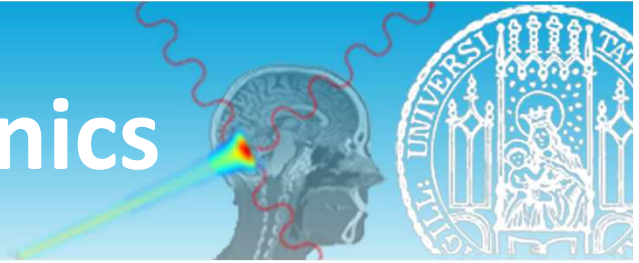


LEX-Photonics: 6J, 20 fs, 5 Hz
 CALA: 60J, 20 fs, 1 Hz
 ~500 mJ, 5 fs, 1 kHz



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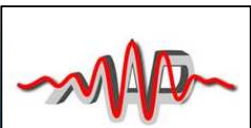
Setup at LEX Photonics



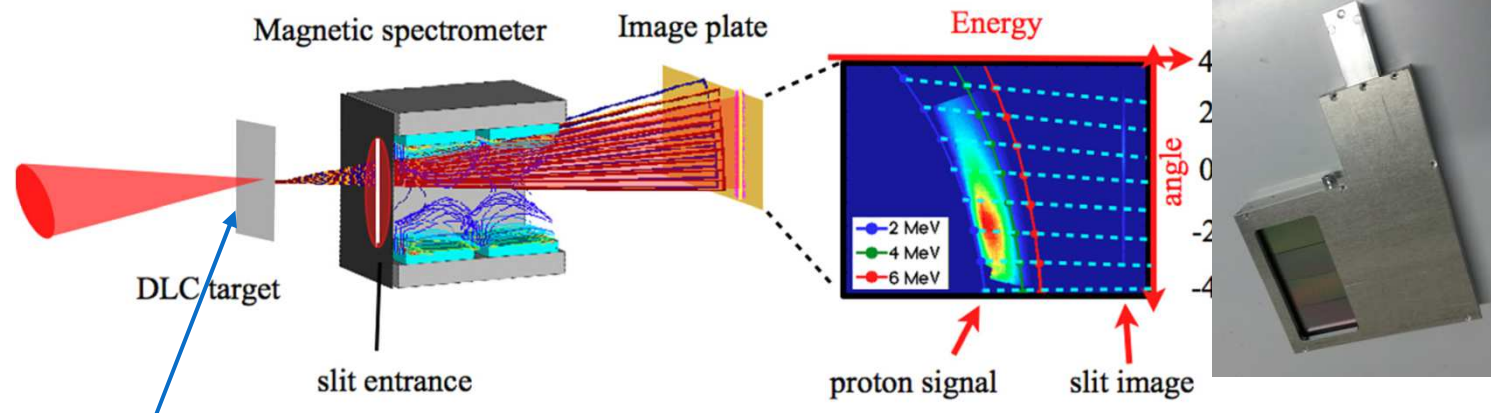
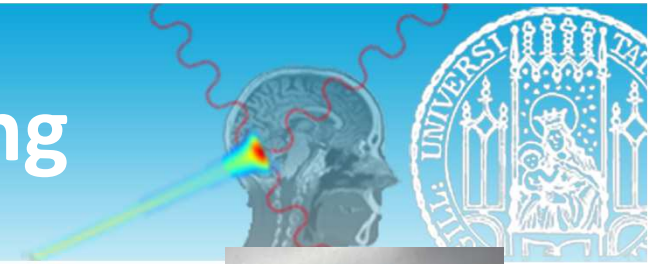
Laser-Beam Delivery

Ion source:
ready to rep-rate

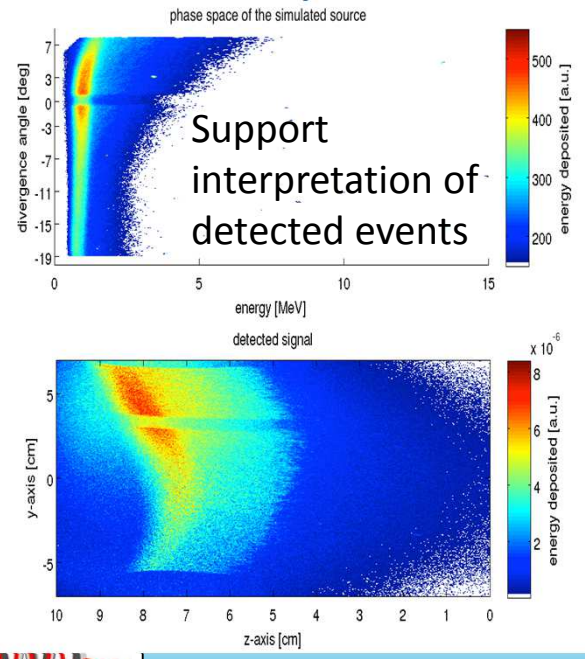
LION target chamber
(prototype for CALA)



Complete Modelling



1
A
p
1

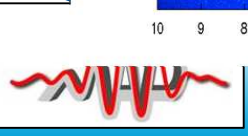


Support interpretation of detected events

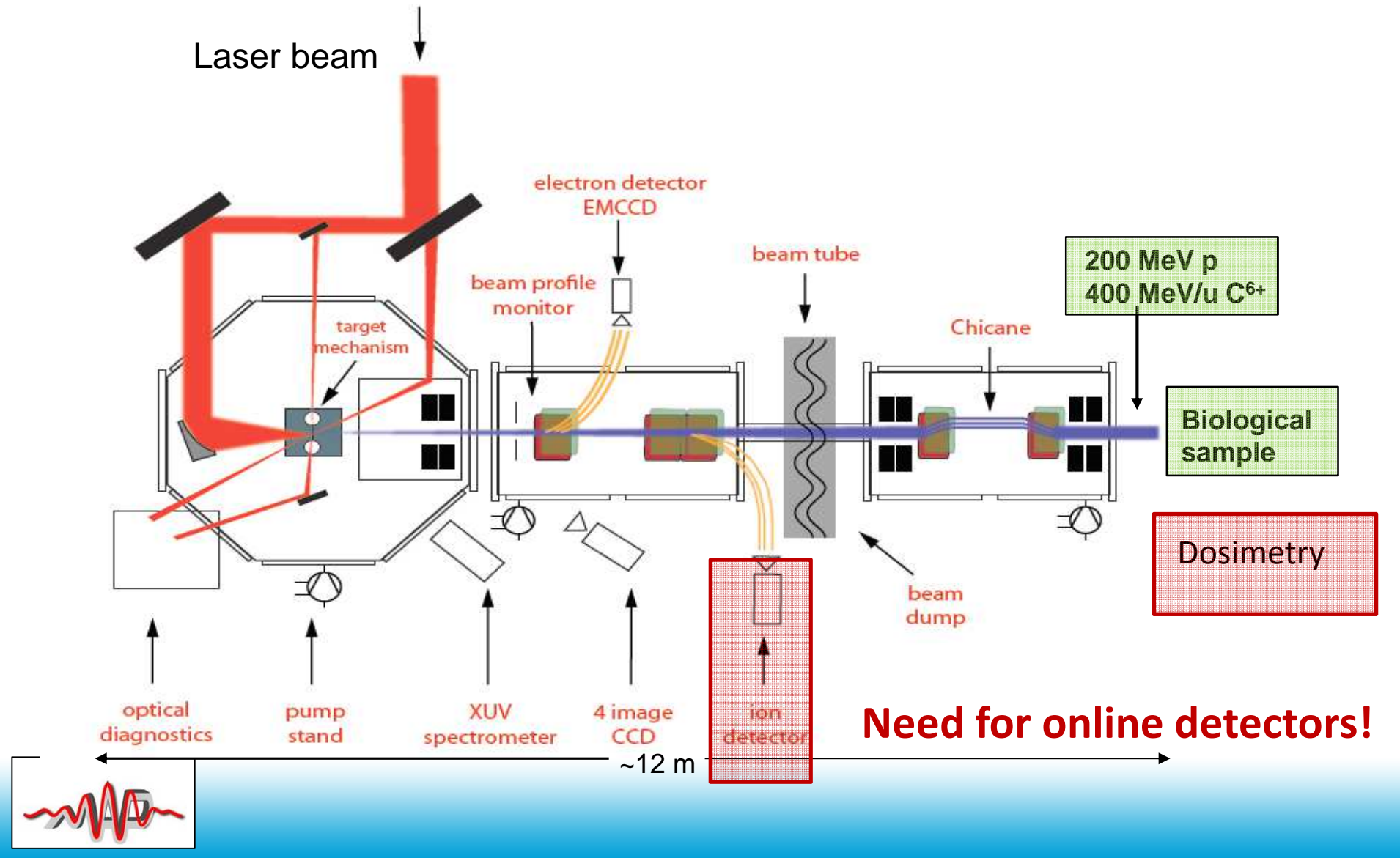
Monte Carlo simulations (Deddes, Parodi)

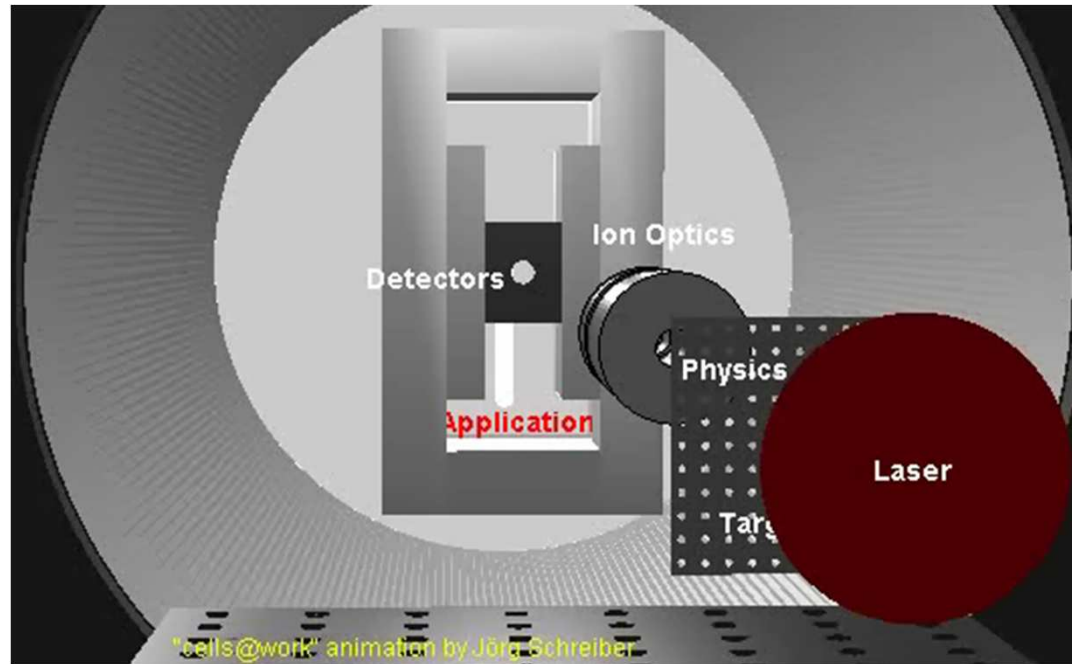
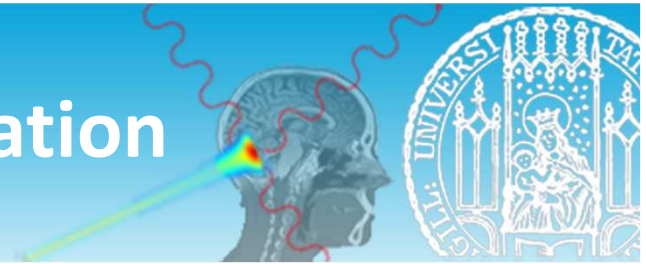
Online Diagnostics

(Reinhardt, Parodi)

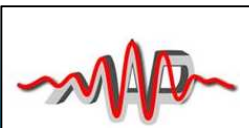


Laser-driven ions: LION





- Laser-Pulses accelerate ions to MeV/u over several μm 's within less than 1 picosecond
- Large ion number in very short time and typically very broad spectrum
- So far mostly single shot/proof of principle experiments



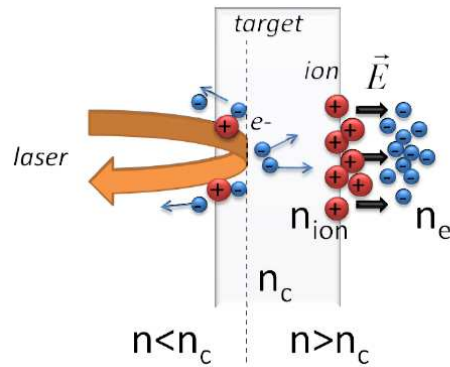


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Laser-driven ion acceleration



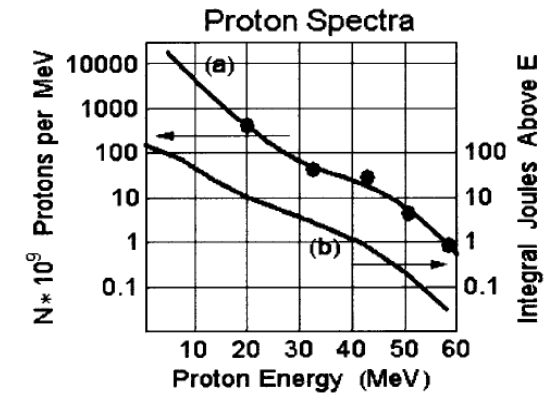
Target Normal Sheath Acceleration (TNSA)



micrometer thick foils

*Maximum proton energies
~ 60 MeV*

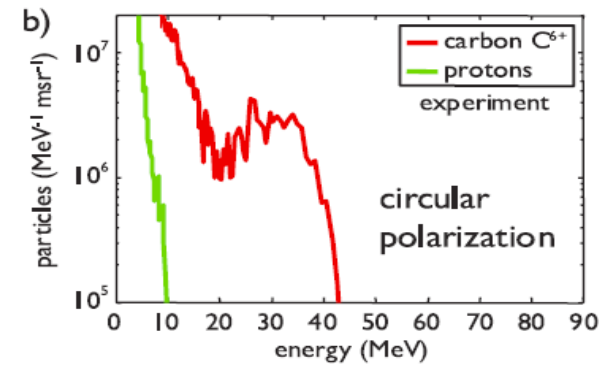
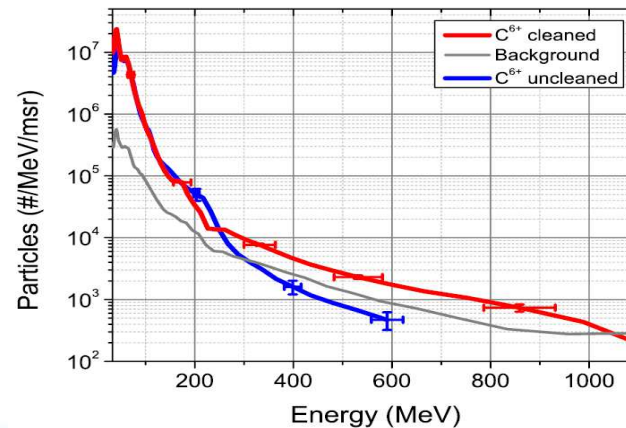
Snively, PRL (2000)



Radiation Pressure Acceleration (RPA)

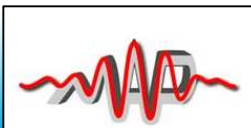
nanometer thick foils

*record carbon energy
> 1 GeV*



A.Henig et al., Phys Rev. Lett. 103, 245003 (2009):
1st experimental demonstration of RPA

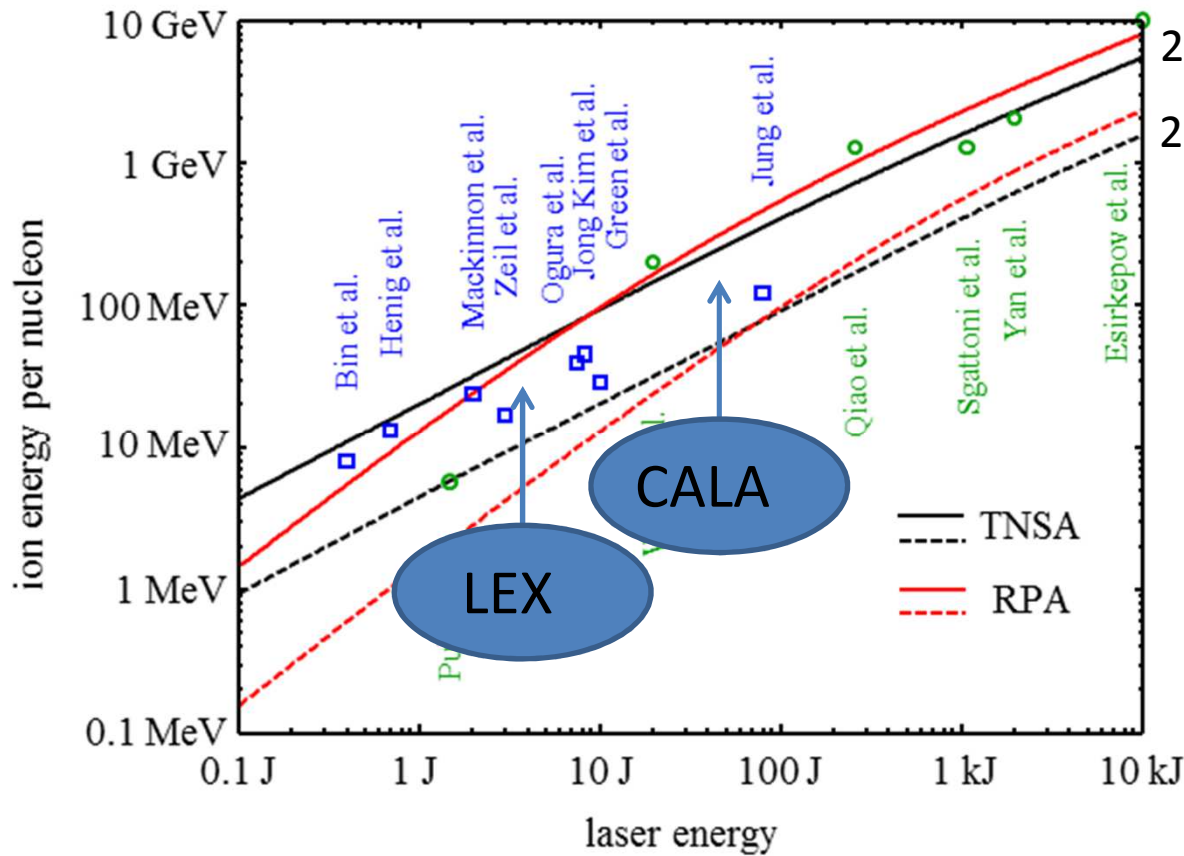
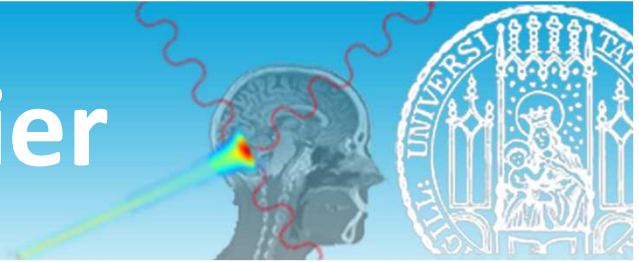
D. Jung et al., Phys. Plasmas 20, 083103 (2013):





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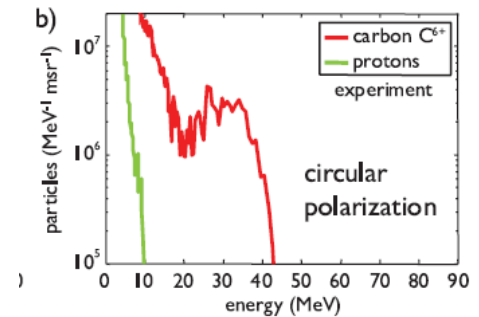
Ion energy frontier



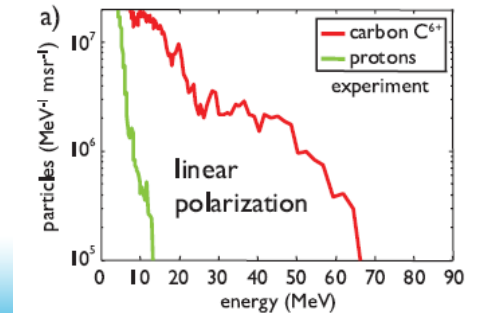
2 μm focus

20 μm focus

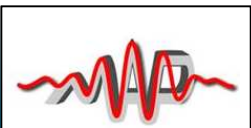
RPA: many ions,
possibly mono-energetic



TNSA: few ions, exponential spectra

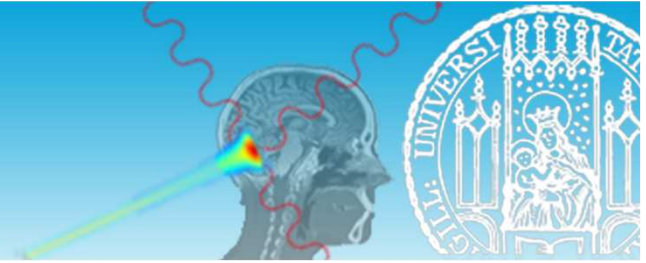


J. Schreiber *et al.*, High Power Laser Science and Eng. 2, e41 (2014)

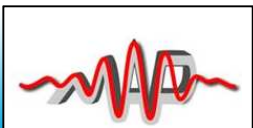




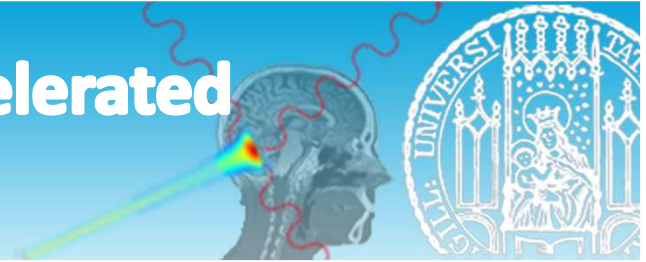
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Detection of laser-accelerated ions

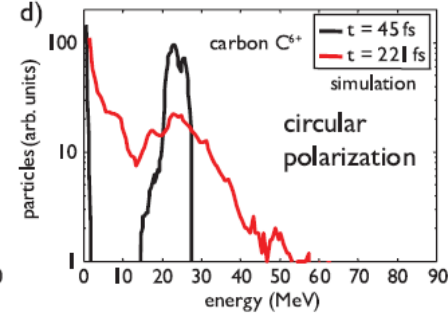
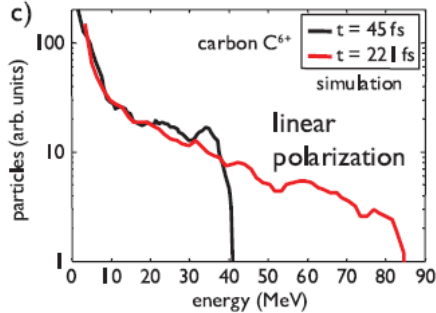
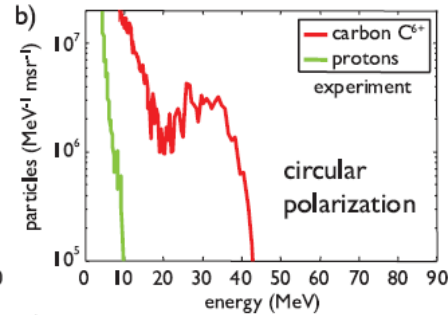
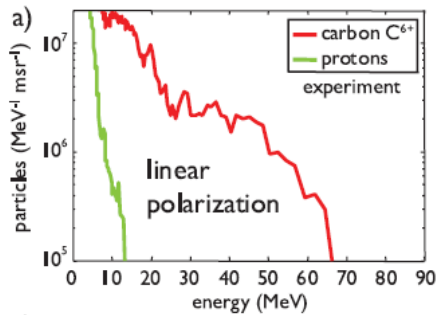


Characteristics of laser-accelerated proton (ion) beams



TNSA

RPA



- **Ultra-short ($\leq ns$) and highly intense ($> 10^7$ ions/cm²) ion pulses**
- EMP presence
- Mixed radiation background
- **Large energy spread** of ions

A. Henig et al, PRL 103 (2009)

→ **Challenge for any electronic online detector but also dosimetry**

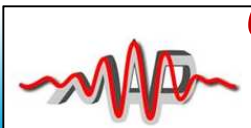
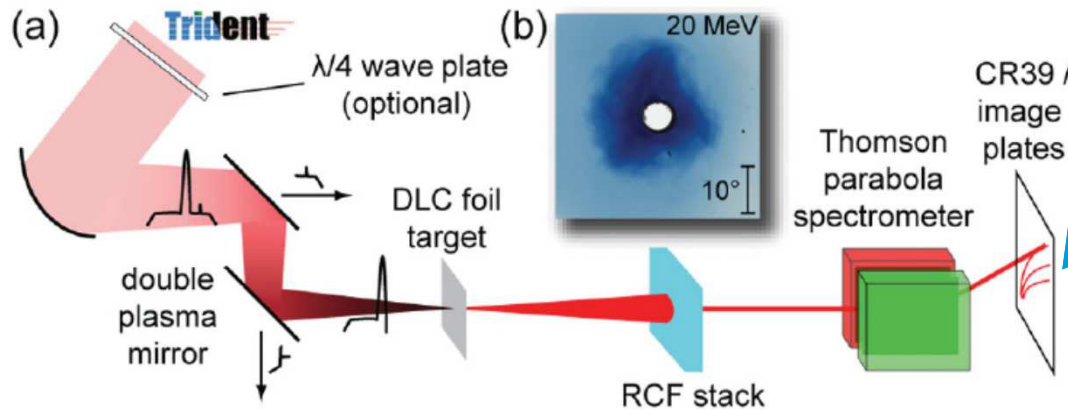
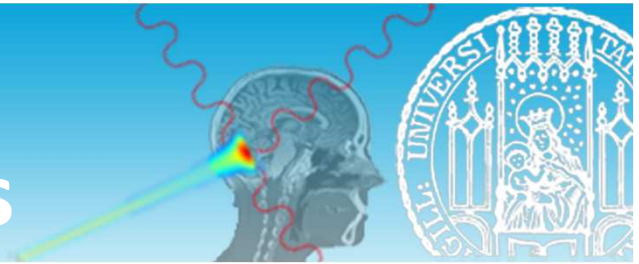




Table 1. Detectors commonly used for laser-driven ion diagnostics and measurements. Sensitivity: L: light, UV: ultraviolet, x: x-rays, e-: electrons; the notation ‘•’ represents ‘sensitive’.

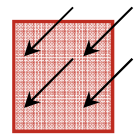
	Spatial resolution	Time resolution	Treatment/ display time	Single- particle sensitivity	Dynamic range (DR)	Sensitivity			
						L	UV, x	e-	
Solid-state nuclear track detectors, e.g. CR-39 (allyl diglycol carbonate), etc	~ a few to a few tens of μm (pit size, depends on the ion kind and energy and etching time)	No	A few hours (etching, scanning, pit counting)	Yes	$\sim 10^2$ – 10^6 (background $\sim 10^2$ – 10^4 cm^{-2} , saturation $\sim 10^6$ – 10^8 cm^{-2})	—	—	—	(1) Sensitive to ions only ^a , single particles
Radiochromic film (RCF)	~3–10 μm (film, scanner)	No	Several minutes (scanning)	No	$\sim 10^2$ – 10^3 (e.g. 10 – 10^4 Gy)	—	•	•	(2) Self-developing
Imaging plate	Sub-100 μm (scanner)	No	Several minutes (scanning)	No	$\sim 10^5$	—	•	•	(3) Reusable, high DR
Activation	Sub-mm (contact radiography)	No	Tens of minutes–a few hours (decay time)	No	Very high ($> 10^5$)	—	—	—	(4) Very high DR
Micro-channel plate (MCP)+phosphor screen + CCD	~several 10 s of μm (imaging system)	~a few 100 ps (MCP gate time)	~a few seconds (CCD readout)	Yes	$\sim 10^3$	—	•	•	(5) Online, single particles
Scintillator + gated I-CCD or EM-CCD	~several 100 μm (multiple scattering, imaging system)	~a few 100 ps (scintillation time)	~a few seconds (CCD readout)	No	$\sim 10^3$	•	•	•	(6) Online, stackable in depth

Detection of laser-accelerated ions



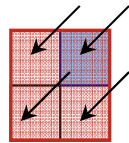
Pixel detector as online detector in Thomson spectrometer:

Advantages of pixel detectors



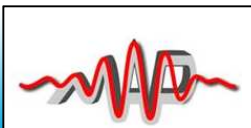
A, N

$$\Phi = \frac{N}{A} = const$$



$$\frac{1}{4} A \rightarrow \frac{1}{4} N$$

- ✓ real time measurement
- ✓ excellent spatial resolution
- ✓ good energy resolution



$$10^8 \text{ particles / cm}^2 = 1 \text{ particle / } \mu\text{m}^2$$



Munich Tandem Accelerator



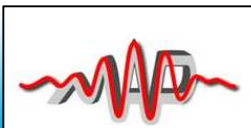
Unique possibilities to test detector response:

- protons: 8- 25 MeV
- 3 irradiation modes:
 - single particle
 - continuous
 - **pulsed: 10^7 protons /cm²/ ns**
→ **similar to laser ion pulse**



Investigated detector systems:

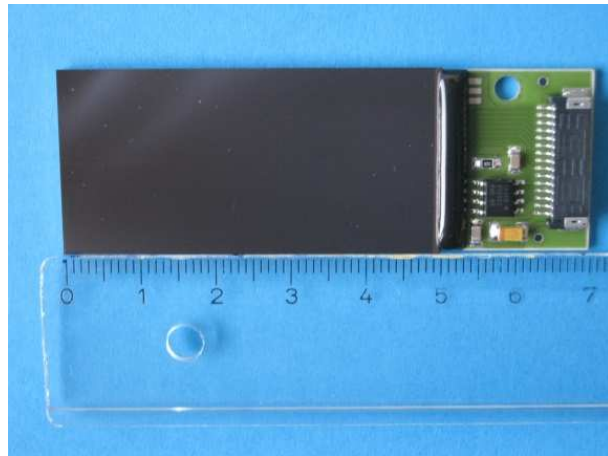
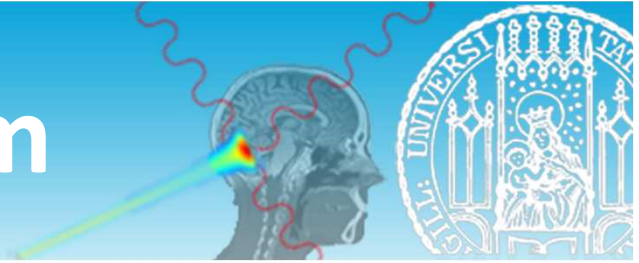
- **Kappa DX-4** (commercial system)
- **Timepix** (scientific & commercial system)
 - collaboration with IAEA CTU Prague
- **RadEye** (commercial system)





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Detector system



RadEye 1 sensor

Silicon pixel detector

512 x 1024 pixel

48 μm pixel pitch

25 x 50 mm^2 sensitive area

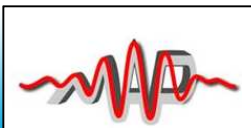


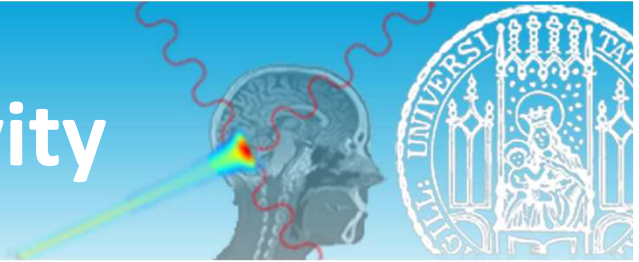
Read-out electronics

Parallel read-out of 4 sensor modules

50 x 100 mm^2 sensitive area

Compact stand-alone system combines computer control and read-out electronics

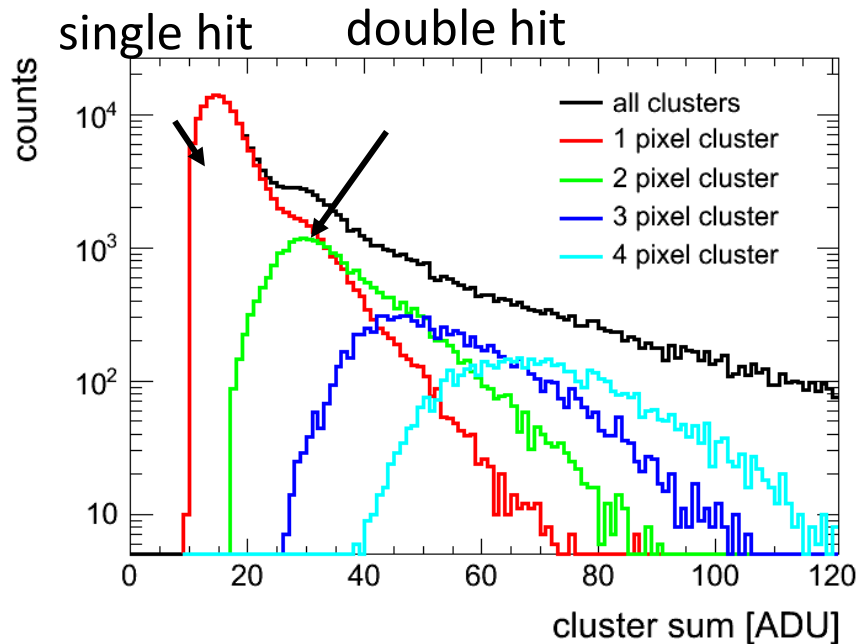




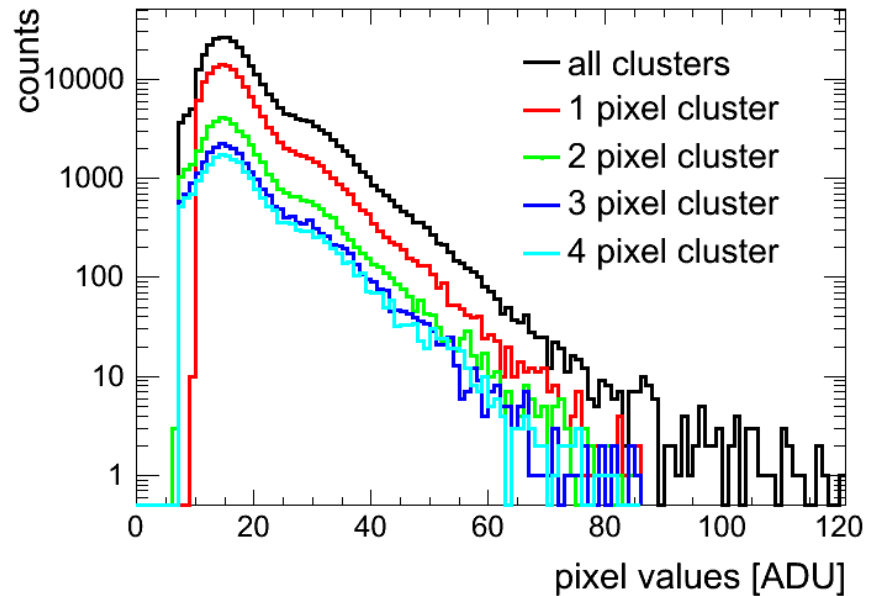
Munich 14 MV Tandem accelerator

- continuous beam
- 15 MeV protons
- $\sim 10^4 \text{p/cm}^2/\text{s}$

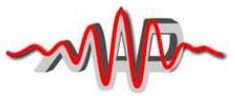
Cluster distribution:



Cluster pixel distribution:



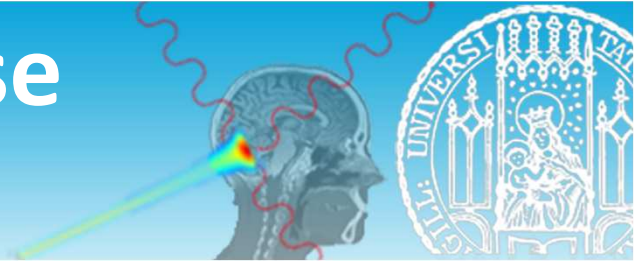
- ✓ *No charge sharing effects observed*
- ✓ *Single and double hits can be distinguished*





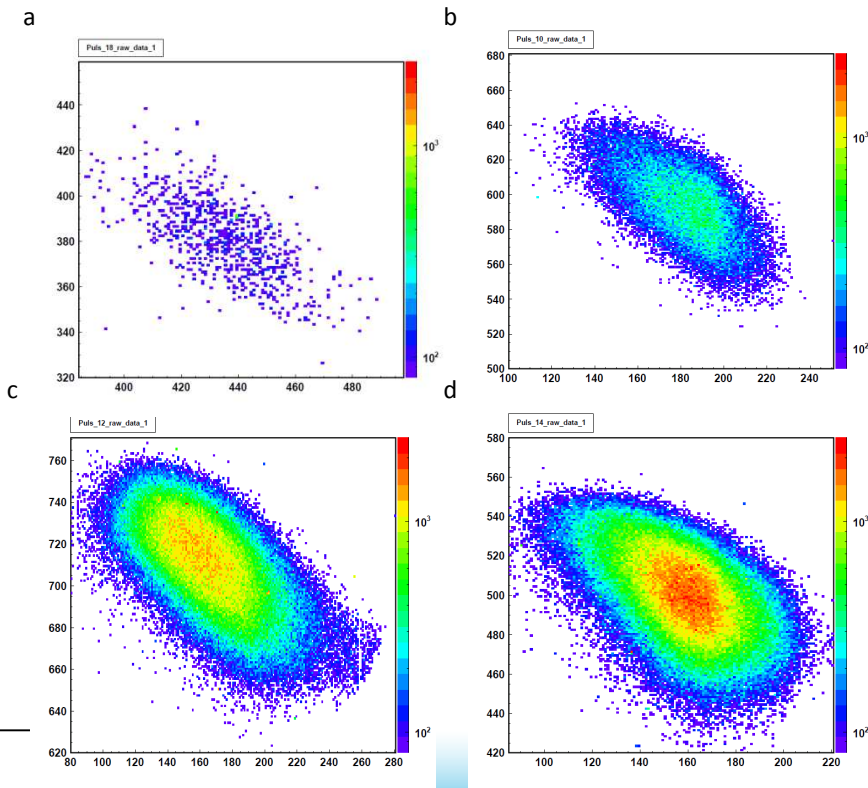
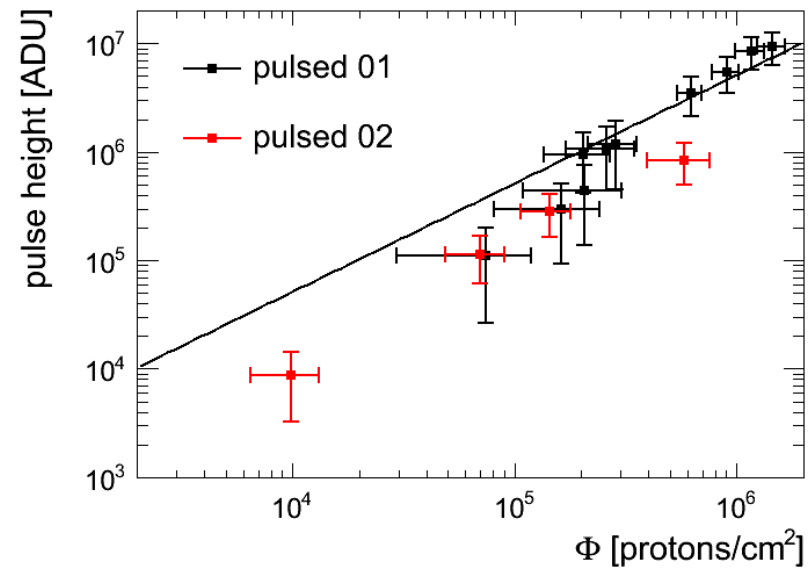
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Intense proton pulse response



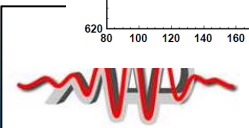
Munich 14 MV Tandem accelerator

- pulsed beam
- 20 MeV protons
- $10^4 - 10^7$ p/cm²/ns



No saturation observed

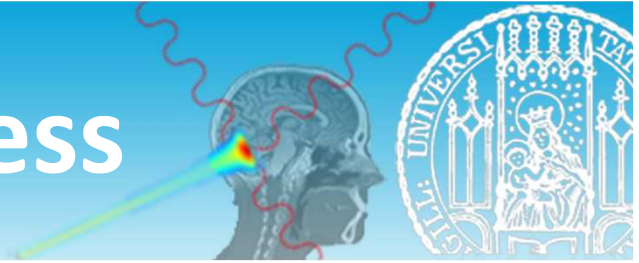
Good agreement to continuous measurements





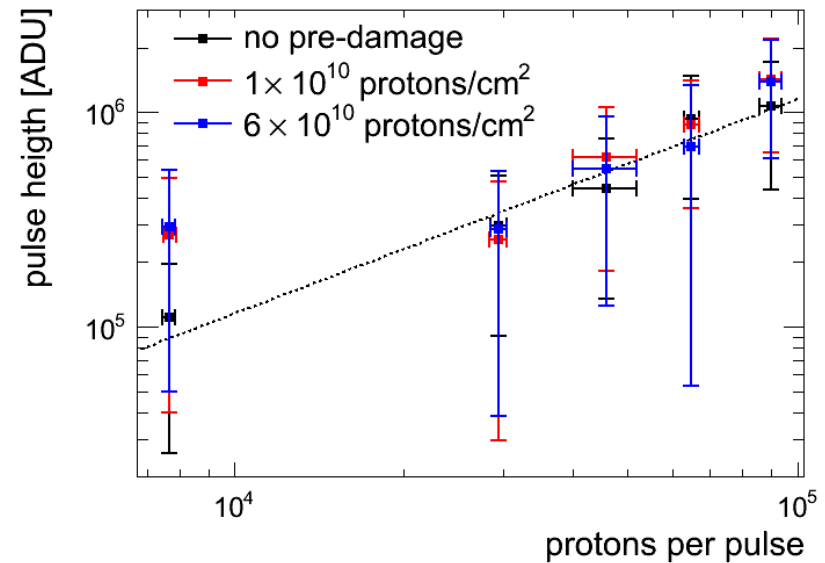
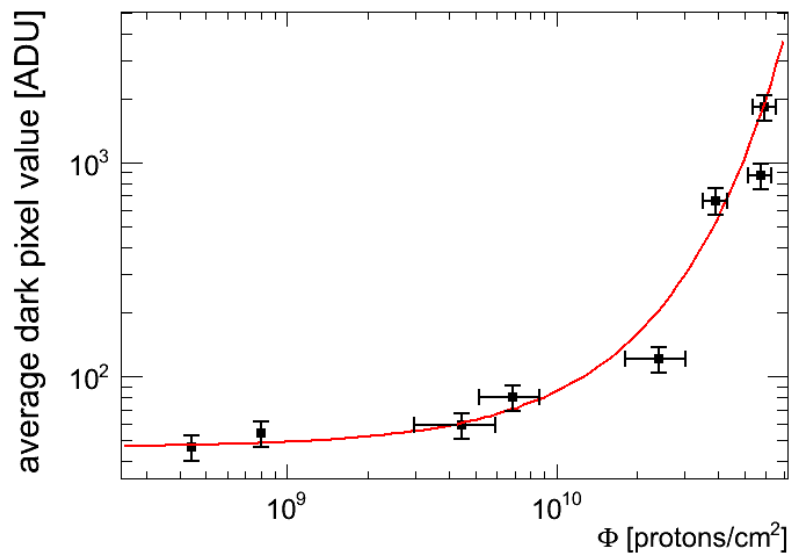
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Radiation hardness



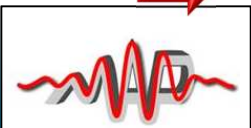
Munich 14 MV Tandem accelerator

- continuous + pulsed beam
- 20 MeV protons
- $\leq 6 \cdot 10^{10}$ p/cm²

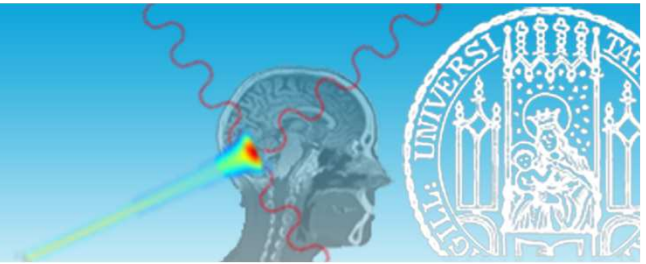


Lifetime \leftrightarrow 90% residual dynamic range

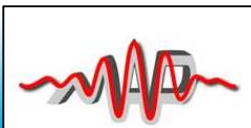
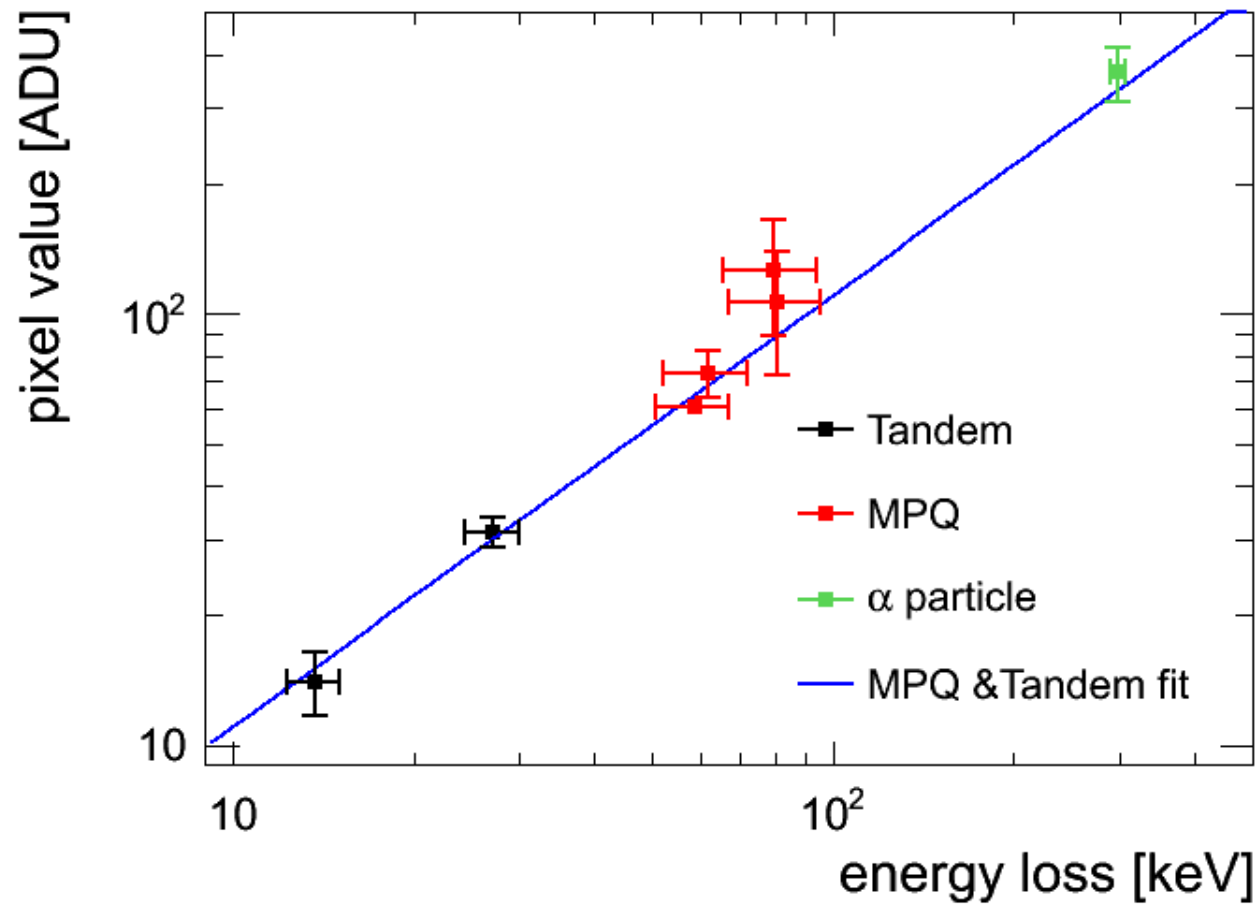
\rightarrow **3000 shots** (20 MeV, 10^7 p/cm²)



System ready for laser-ion-acceleration experiment !



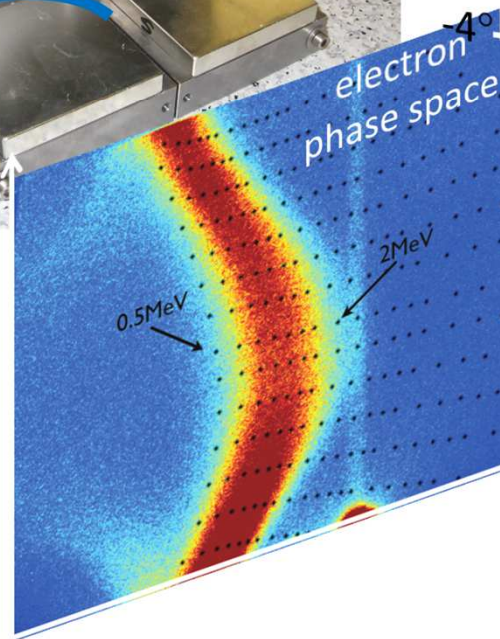
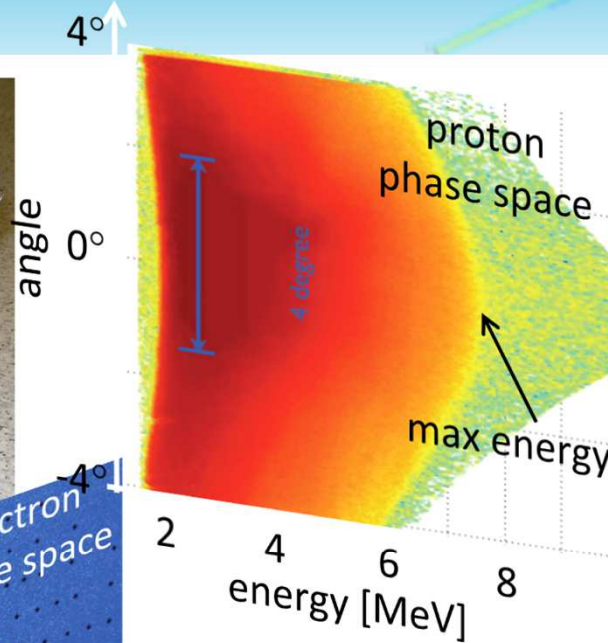
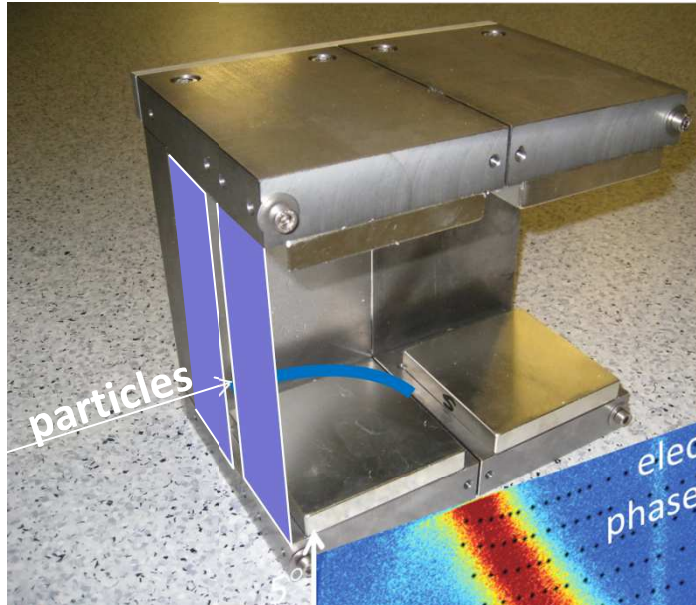
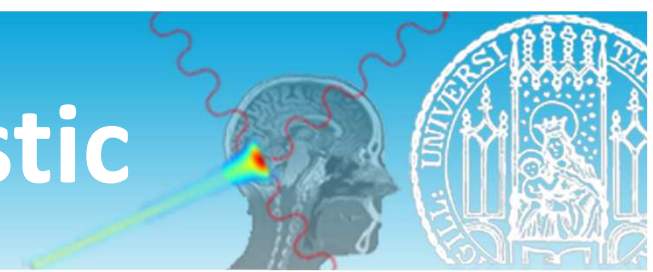
Energy conversion: 1.11 ± 0.09 ADU/keV



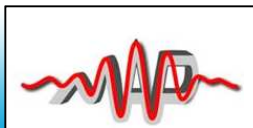


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LEX LION Diagnostic

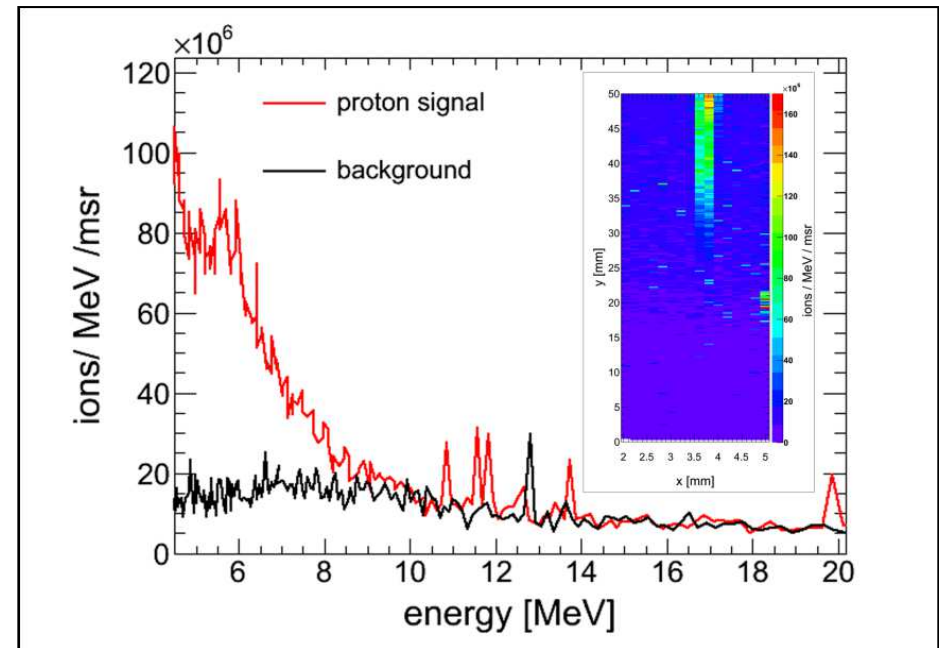
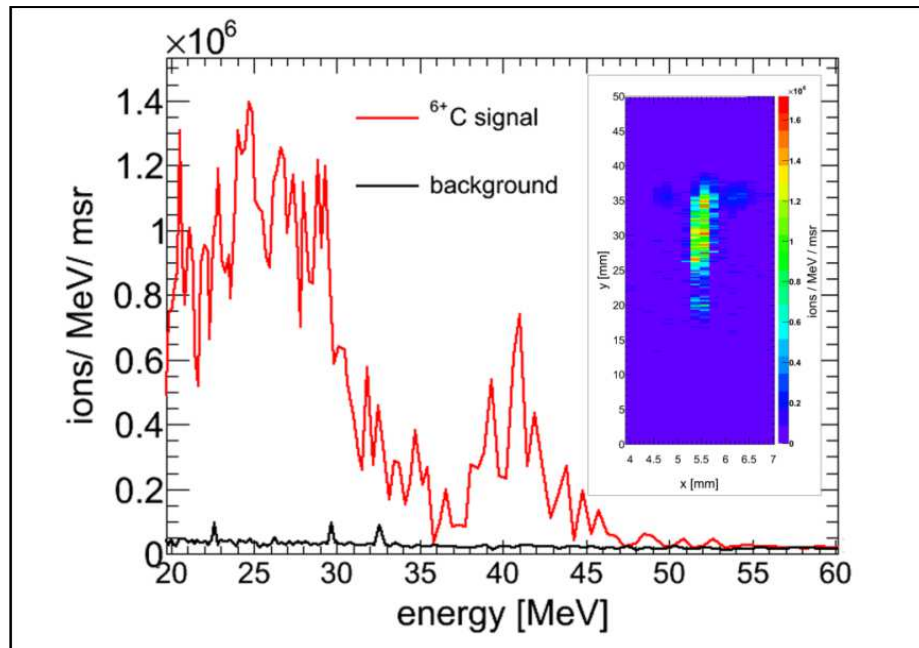


Particle spectrometer combined with RadEye detector

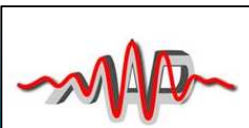




- simultaneous detection of protons and 6^+C -ions
- 10% accuracy in particle number determination possible

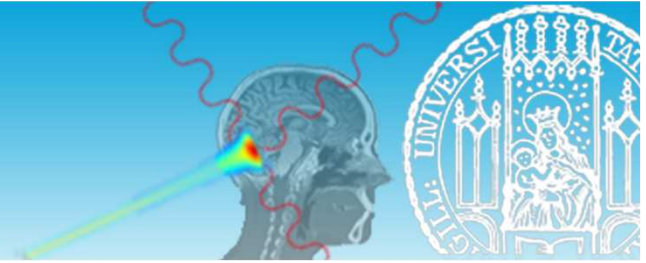


S. Reinhardt *et al.*, Journal of Instrumentation 8, 03008 (2013)

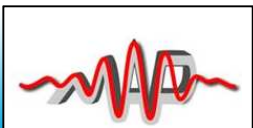




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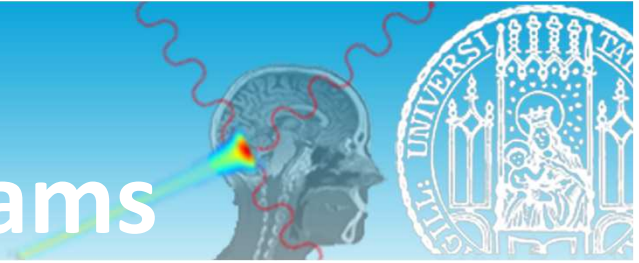


Dosimetry of laser-accelerated ions



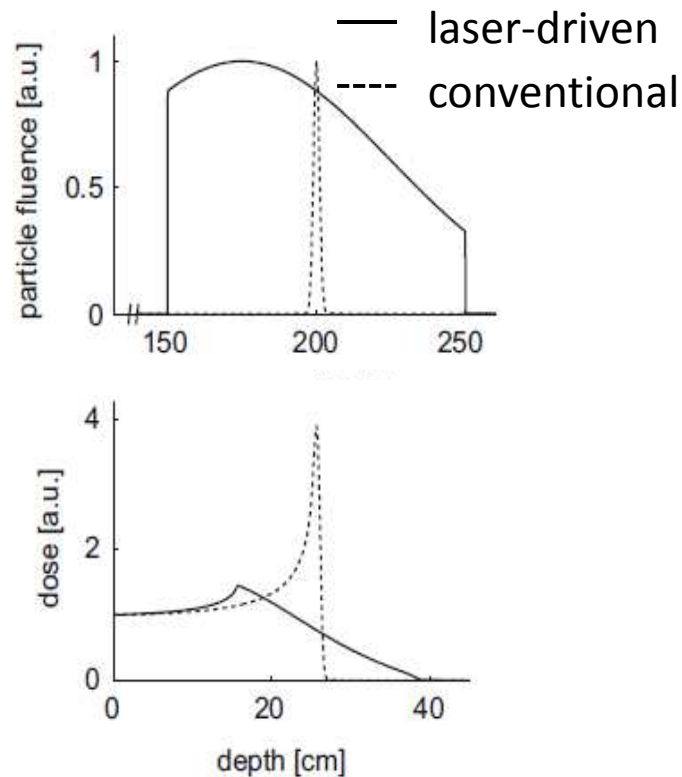


Radiotherapy with laser-driven ion beams

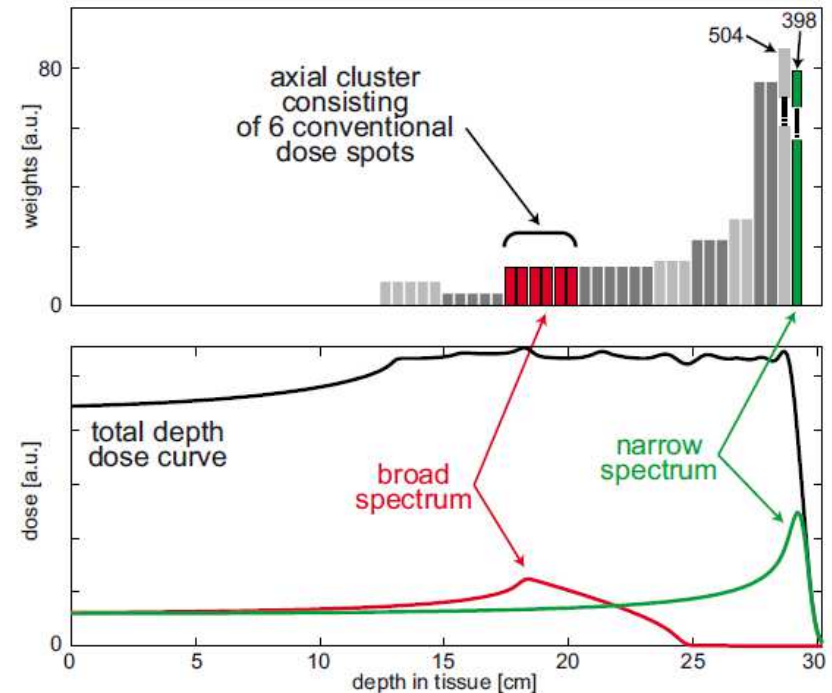
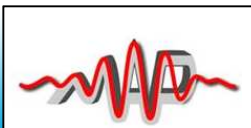


MAP main goal: Radiation therapy with laser-accelerated ions

Main idea:
Exploit broad energy spectrum to generate SOBP



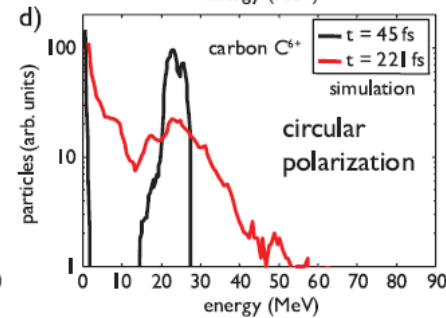
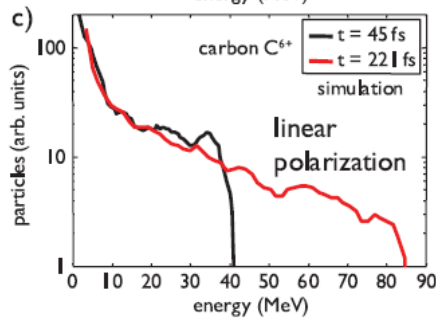
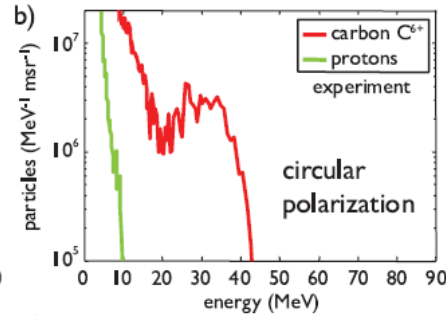
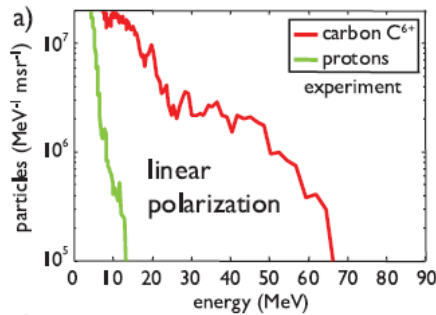
S. Schell et al, Med Phys 37 (2010)





TNSA

RPA

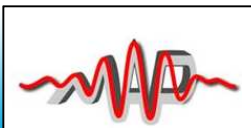


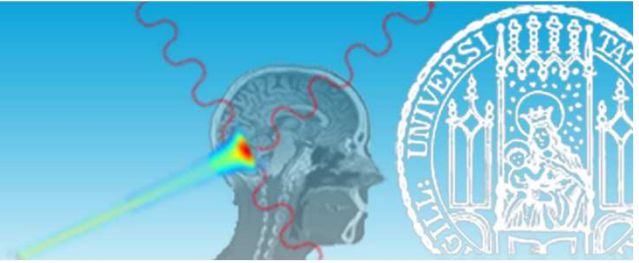
A. Henig et al, PRL 103 (2009)

- **Ultra-short ($\leq ns$) and highly intense ($> 10^7$ ions/cm²) ion pulses**
- EMP presence
- Mixed radiation background
- **Large energy spread** of ions

→ **Challenge for dosimetric measurements**

→ **Investigation of biological response required**





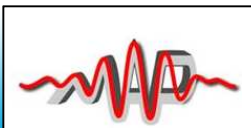
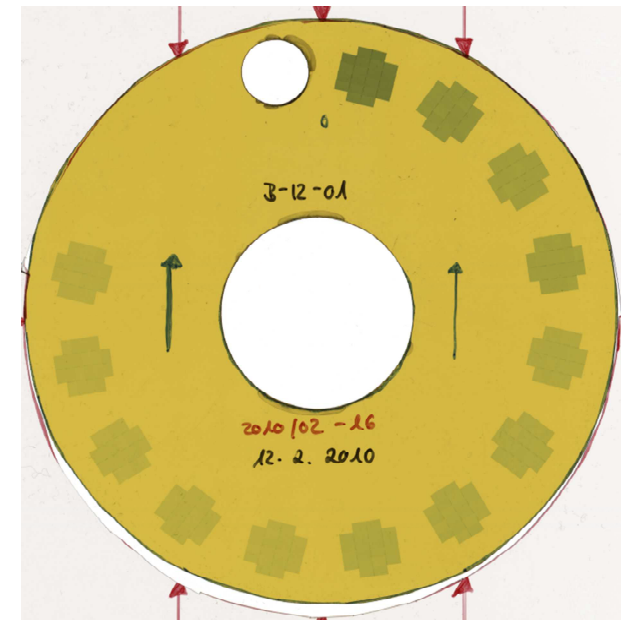
MAP: Radiation therapy with laser-accelerated ions
→ Bio-medical studies at LEX/CALA → **Dosimetry**

➤ **Special dosimetric application**

- high pulse dose rate ($> \text{Gy/ns}$)
- Low energies
(protons $< 20 - 30 \text{ MeV @ LEX}$)

➤ Development of **film dosimetry** protocol:
Radiochromic EBT2/EBT3

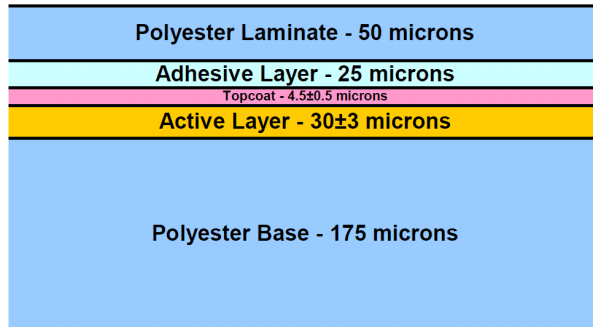
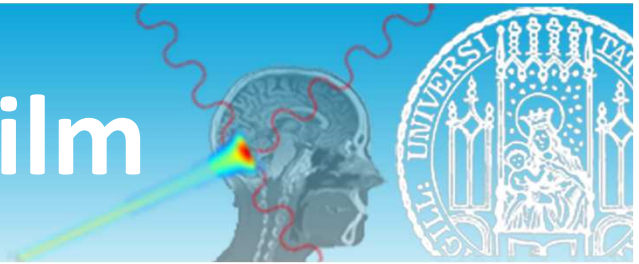
➤ Application in cell and mouse tumour
irradiation





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Gafchromic EBT film



EBT2

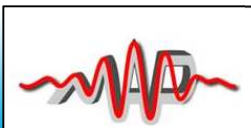
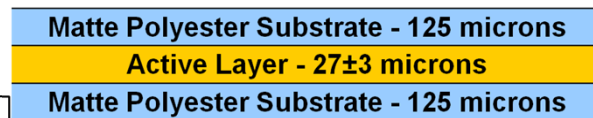
same active layer



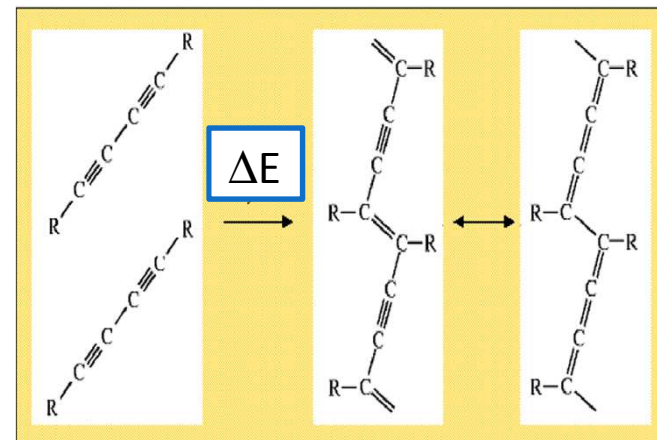
*Lithium pentacos-
10,12-diynoate (LiPCDA)*

EBT3

but symmetric
configuration

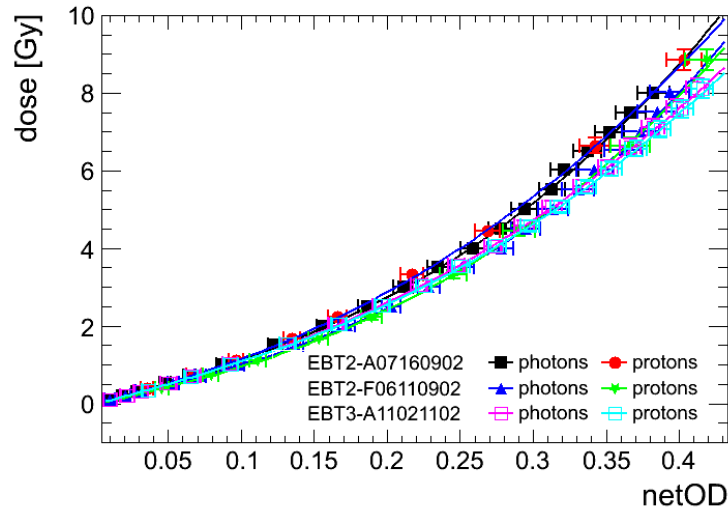
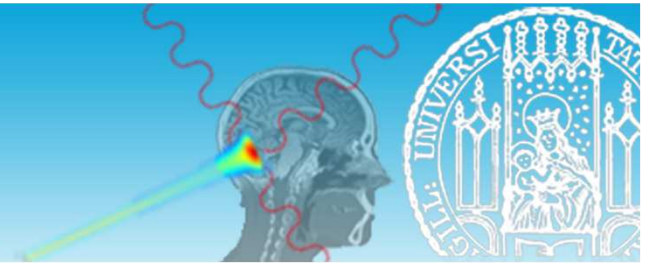


- ✓ Dose range: **0.2 - 40 Gy**
- ✓ Water-equivalent
- ✓ Sub-mm spatial resolution
- ✓ 2D read-out by RGB flatbed scanner
- ✓ Self-developing by polymerization





Film dosimetry in proton beams



EBT2 vs. EBT3:

No general response difference

- No particle type dependence
 - difference photons-protons < 3.0 %
- Intra-batch variations < 2.5 %
- Batch-to-batch variations < 11.5 %

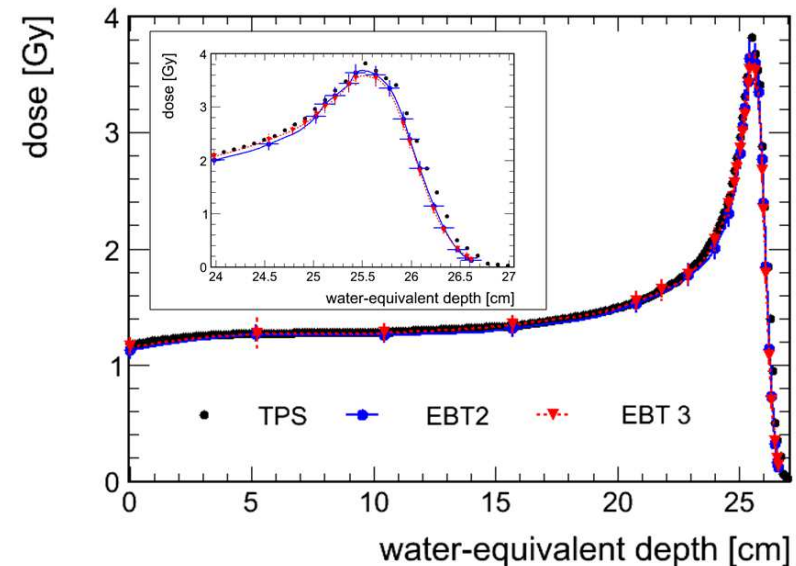
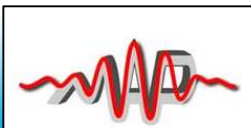
Problem:

- dose under-response in Bragg Peak region (high LET)
- already known from other radio-chromic films

S. Reinhardt et al, Med. Phys. 39 (2012)

S. Reinhardt et al, Rad Env Biophys (2015)

DOI 10.1007/s00411-014-0581-2





Mouse tumour irradiation at the MLL Tandem accelerator

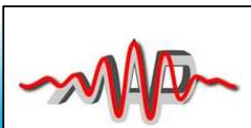
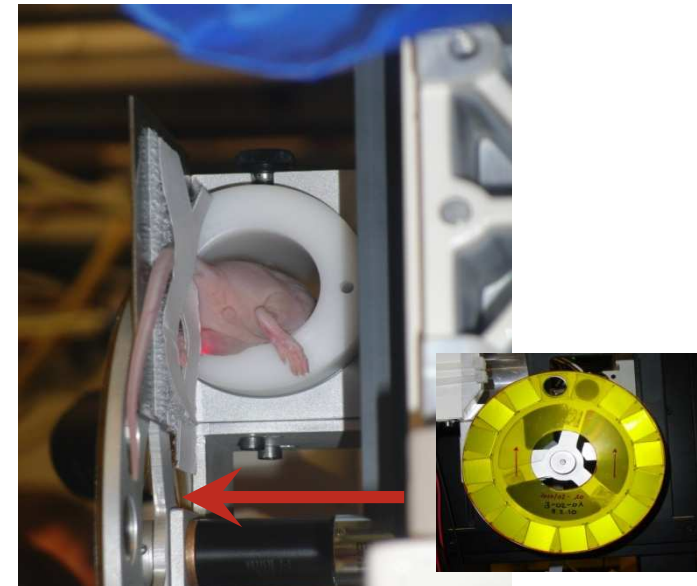


- RBE Difference due to irradiation mode?
(continuous vs. pulsed)

- **23 MeV protons:**
 < 6 mm range in water

- **spatial dose distribution** required
 → **Gafchromic EBT2 - films** for fluence measurements

- **PTV dose: 20 Gy**, using 5 fluences
 → dose range on film: **0.54 Gy - 4.00 Gy**





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Mouse tumour irradiation at the MLL Tandem accelerator



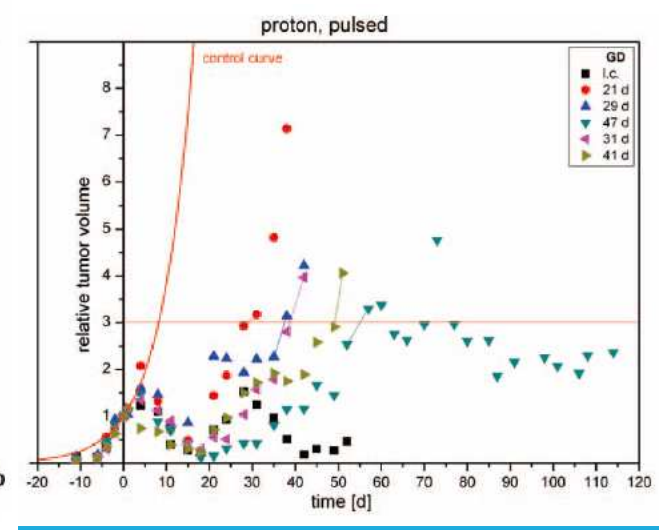
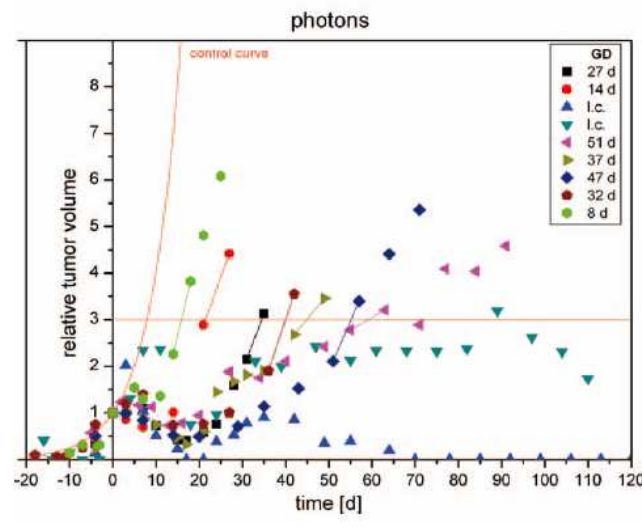
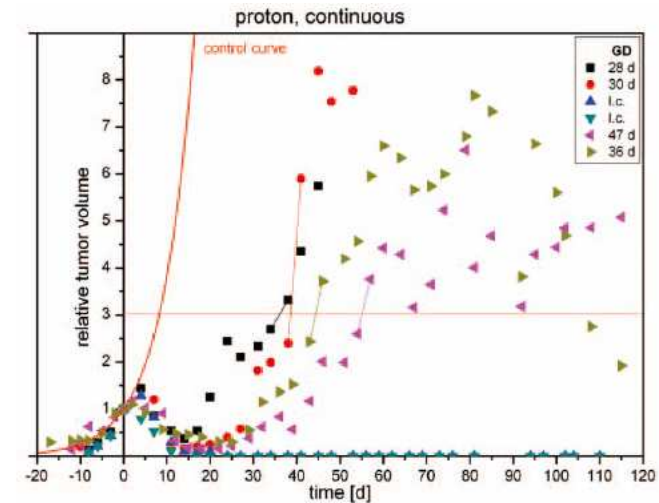
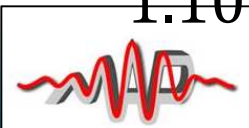
O. Zlobinskaya et al, Rad Res 181(2014)

Our *in vitro* and *in vivo* studies showed no evidence of a substantially different radiobiology associated with the ultra-high dose rate that characterizes protons generated from advanced laser technology.

We conclude that dose prescription for pulsed protons can be based on established therapeutic concepts for protons. However, one should bear in mind that differences in the RBE values smaller than 10% cannot be excluded yet and should be accounted for in dose constraints for organs at risk.

RBE values:

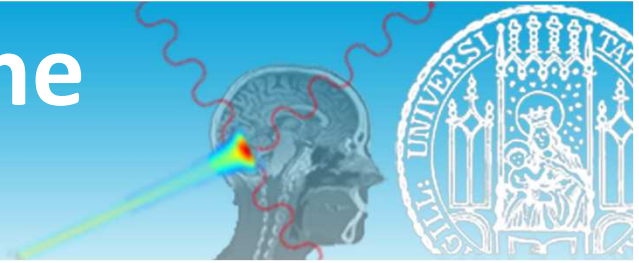
- pulsed beam
 1.22 ± 0.19
- continuous beam
 1.10 ± 0.18



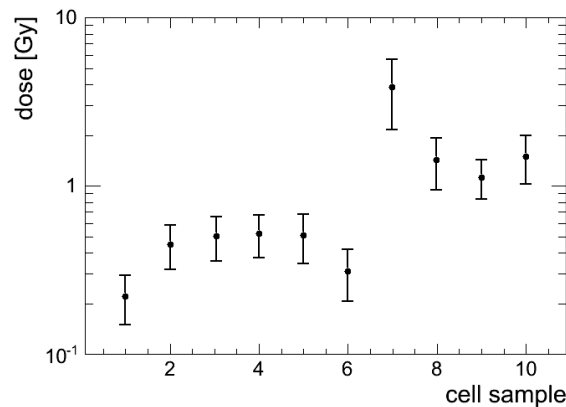
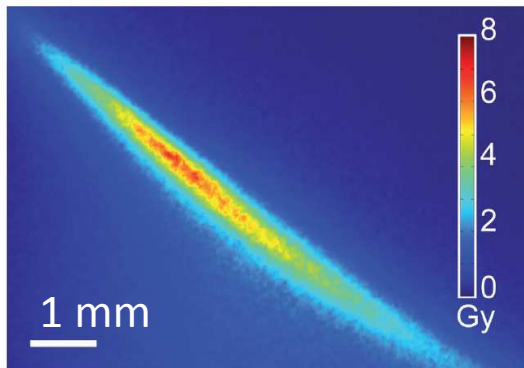
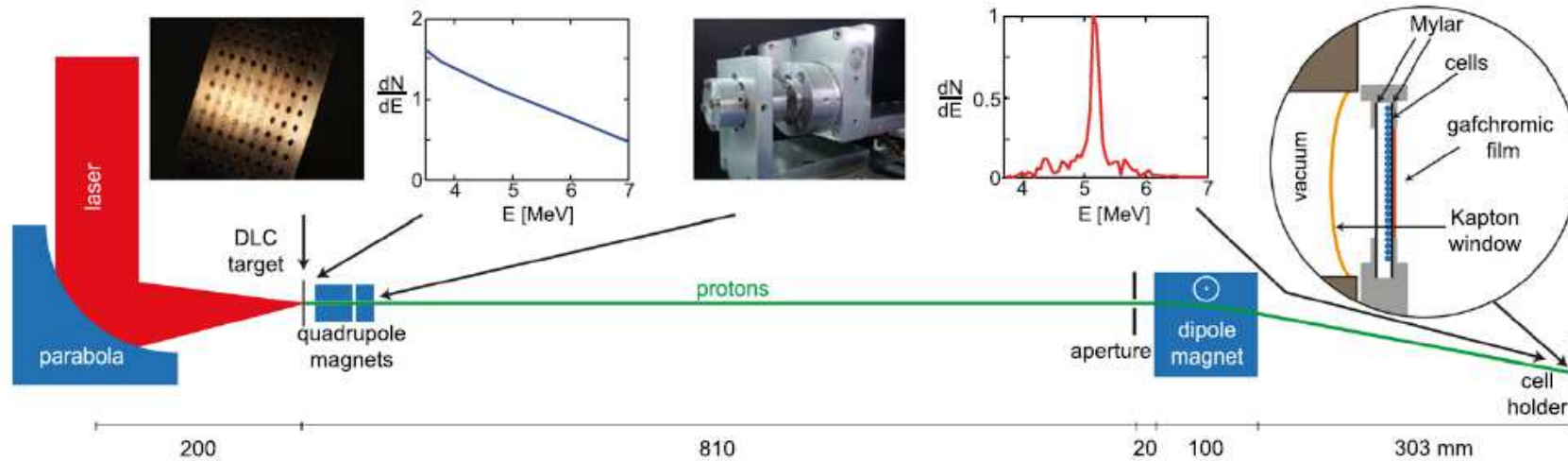


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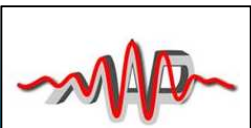
Cell irradiation at the MPQ ATLAS laser



J. Bin et al, APL 101 (2012)



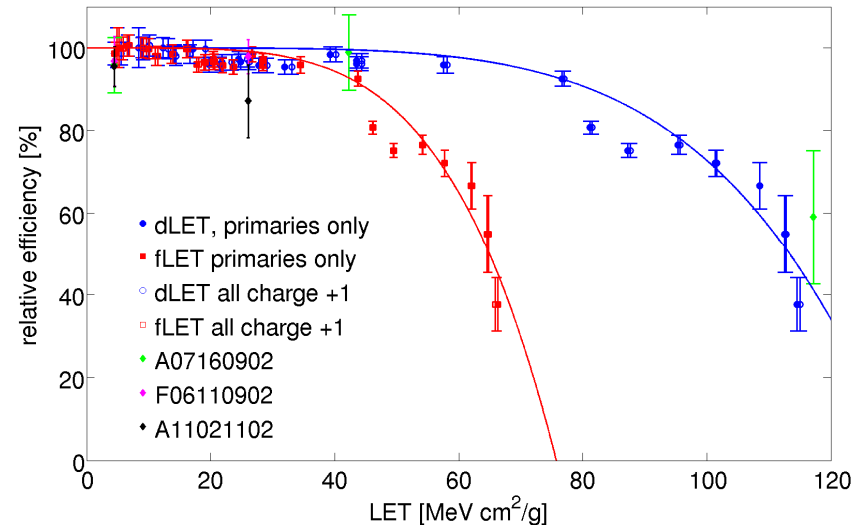
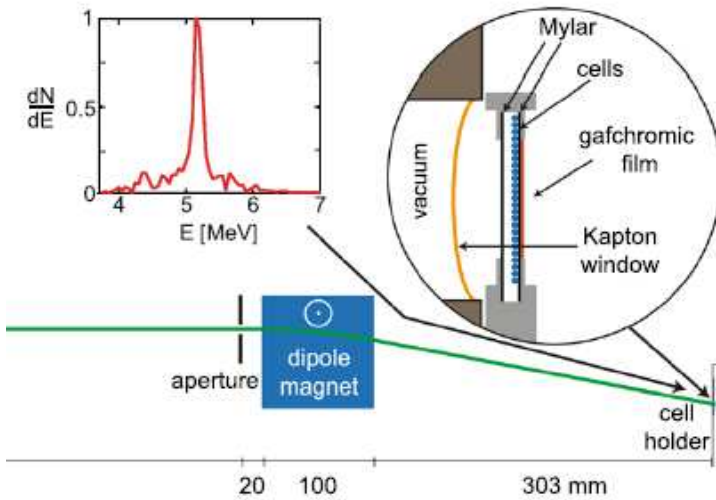
Delivery of single shot doses up to 7 Gy to living cells (HeLa)





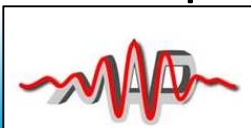
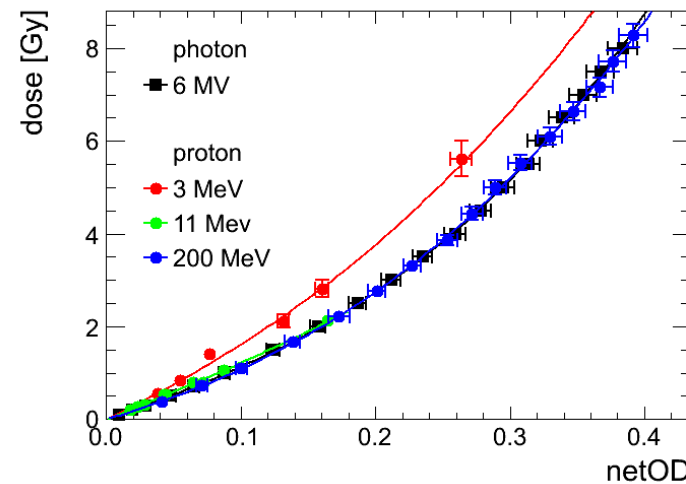
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Cell irradiation at the MPQ ATLAS laser



MPQ cell irradiation experiment 3.1 MeV protons @ active film layer

- considerable LET dependence
- low energy calibration required



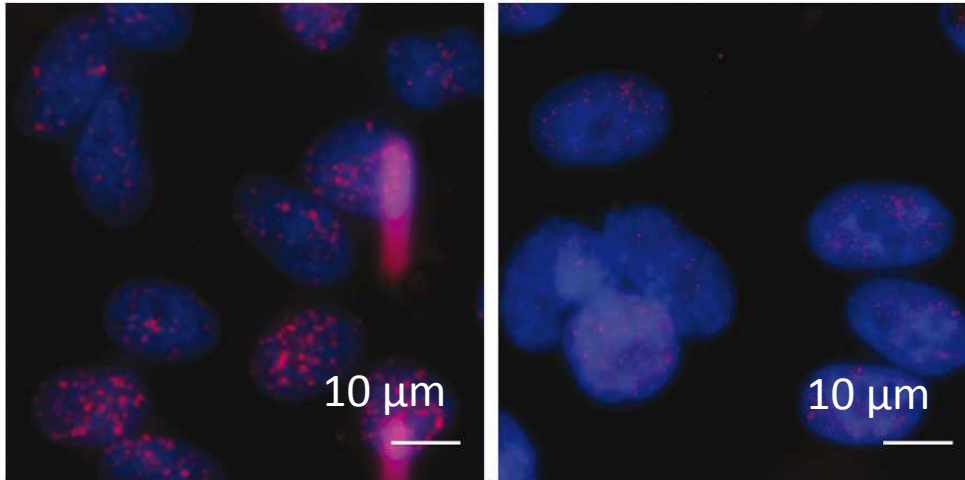
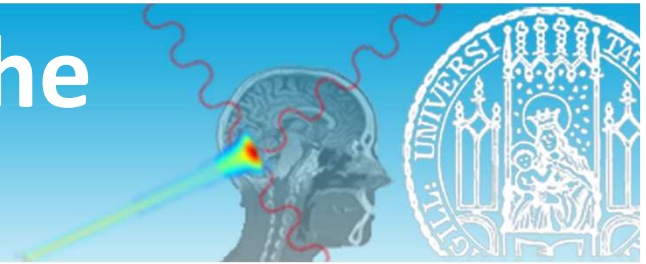
S. Reinhardt et al, Rad. Env. Biophys. (2015)

10.1007/s00114-014-0501-0



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Cell irradiation at the MPQ ATLAS laser



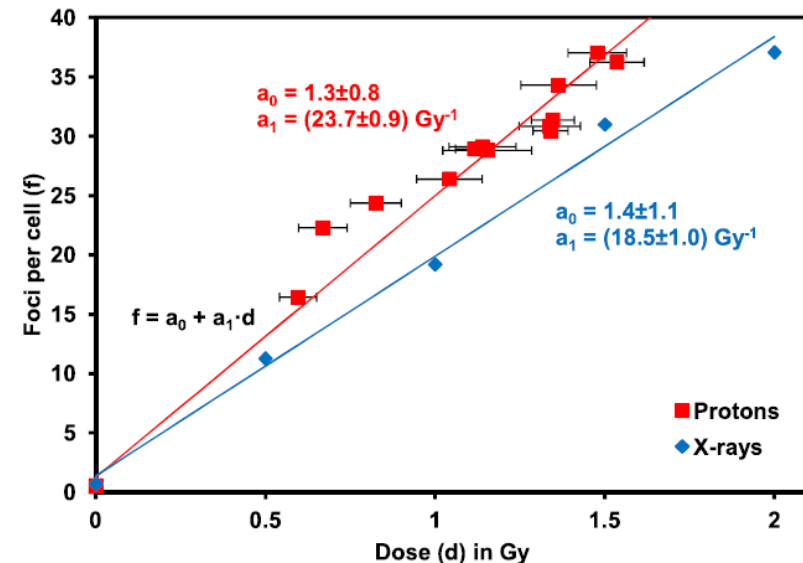
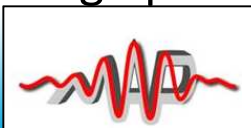
J. Bin et al, APL 101 (2012)

FIG. 5. Initial DNA damage in HeLa cells. (a) Sample exposed to a mean dose of 1.0 Gy and (b) corresponding unirradiated control. Foci of γ -H2AX (red) and cell nuclei (blue) are shown (3D microscopy, maximum intensity projections, background correction, contrast enhanced). The red vertical bars in (a) are part of the grid used for spatial registration (Fig. 4). Horizontal scale bars, 10 μ m.

Irradiation of HeLa cells:

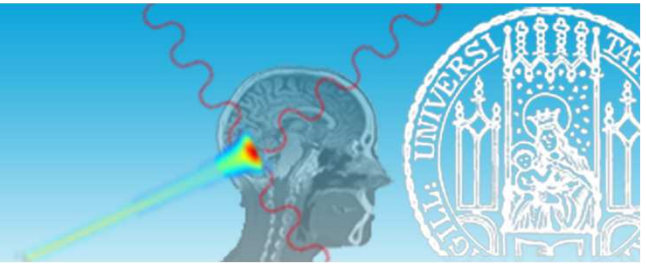
RBE for induction of repair foci
 1.3 ± 0.3

No new radiobiological effects by single
high pulse dose rate $\sim 10^9 \frac{\text{Gy}}{\text{s}}$





Conclusion and Outlook



Gafchromic films

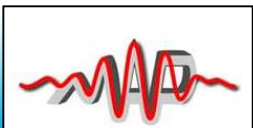
- EBT3 and EBT2 show similar performance
- Film dosimetry in low energy proton beams possible

RadEye-Detector

- ***Single proton sensitivity***
- Limited energy resolution
- Sufficient radiation hardness
- ***Linear pulse dose response up to 10^7 p/cm²***

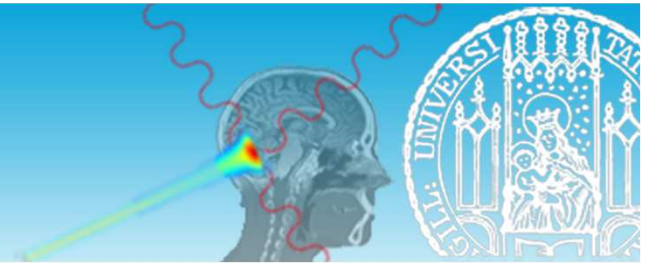
Outlook

- *RadEye in routine use for laser accelerated ion detection in LEX*
- *New detector developments required for ion detection in CALA*
- *Online dosimetry required for future experiments in LEX/CALA*





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A. Alhazmi, W. Assmann, R. Berger, G. Dedes, A. Mayr, S. Reinhardt, J. Schreiber, K. Parodi

Thank You !

