

Small animal micro-CT & mixed radiation field characterization in ion beam radiotherapy[#] and in outer space^{*} with pixel detectors Timepix



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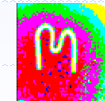


@ PhD student



Research performed in frame of the CERN Medipix Collaboration

^{*} Project funded by the European Space Agency





Colleagues/Co-authors/Acknowledgements

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Small animal μCT

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Basic Research

Astroparticle & non-accelerator physics

- Neutrino physics (NEMO3/SuperNEMO, TGV)
- Cosmic rays (CZELTA)
- Dark matter (PICASSO)

ATLAS at LHC

- SCT detection modules
- Neutron shielding
- Medipix radiation monitoring
- Higgs boson physics

Nuclear spectroscopy

- Fission fragment spectroscopy
- Laser induced nuclear excitation
- Ultra cold neutrons

Applied Research

Radiation imaging

- Medipix pixel detectors: SW, HW
- X-ray radiography and tomography
- Charged particle & neutron imaging
- Biomedical imaging
- Material science and defectoscopy

R&D of semiconductor detectors

- 3D and semi-3D detectors
- Thermal neutron detectors
- Room-temperature detectors
- Instrumentation for detector testing

Applied spectrometry

- Material analysis (CINAA, XRF, Radon)
- Particle tracking and spectroscopy
- Space: (gamma, neutron, micro-sensor, SATRAM payload)

Fundamental Experiments in the Physics of the Microworld

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- [NSS MIC IEEE Conference](#)
Seattle, USA
8-15 Nov 2014
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27 Oct - 2 Nov 2013
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Paris
23-27 June 2013

2.5 MeV VdG accelerator at IEAP CTU in Prague

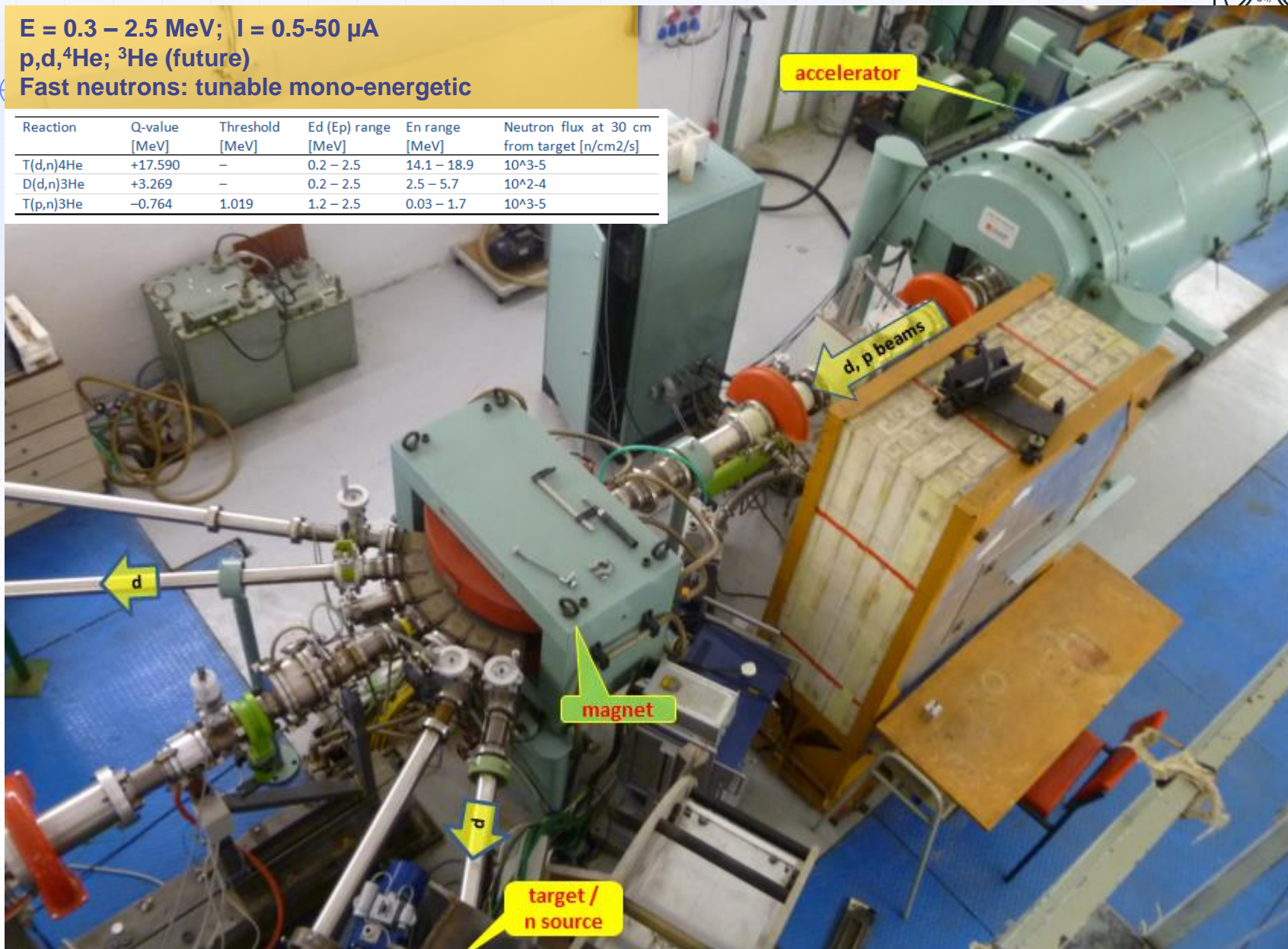


$E = 0.3 - 2.5 \text{ MeV}$; $I = 0.5-50 \mu\text{A}$

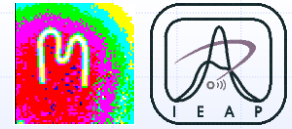
$p, d, {}^4\text{He}$; ${}^3\text{He}$ (future)

Fast neutrons: tunable mono-energetic

Reaction	Q-value [MeV]	Threshold [MeV]	E_d (E_p) range [MeV]	E_n range [MeV]	Neutron flux at 30 cm from target [$\text{n}/\text{cm}^2/\text{s}$]
$T(d,n){}^4\text{He}$	+17.590	-	0.2 - 2.5	14.1 - 18.9	10^{3-5}
$D(d,n){}^3\text{He}$	+3.269	-	0.2 - 2.5	2.5 - 5.7	10^{2-4}
$T(p,n){}^3\text{He}$	-0.764	1.019	1.2 - 2.5	0.03 - 1.7	10^{3-5}



Small animal micro-CT& mixed radiation field characterization in ion beam radiotherapy[#] and in outer space^{*} with **pixel detectors Timepix**



Hybrid semiconductor pixel detectors Medipix/Timepix

Per-pixel signal readout electronics

Institute of Experimental and Applied Physics
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Hybrid architecture: sensor is bump-bonded to the readout ASIC chip. Different semiconductor sensors can be used:

- material (Si, CdTe, GaAs)
- thickness (e.g. 50, 300, 500, 700, 1000, 1500 μm).

- high granularity
- sub-pixel & sub- μm spatial resolution
- per-pixel signal processing (13-bit/px)



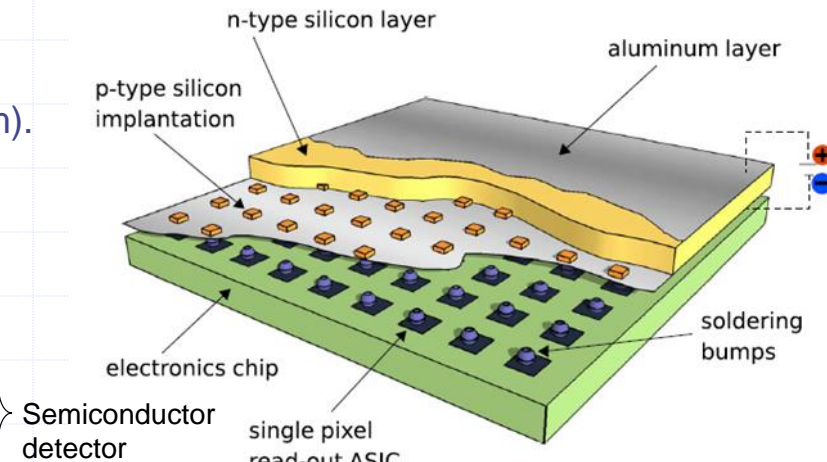
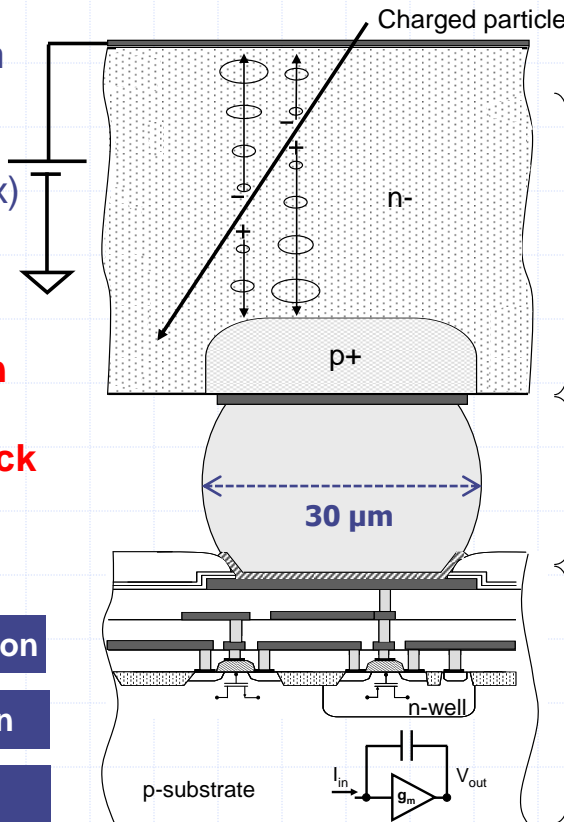
- dark-current free **noiseless detection**
- threshold ~ 4 keV
- **single-particle track visualization**



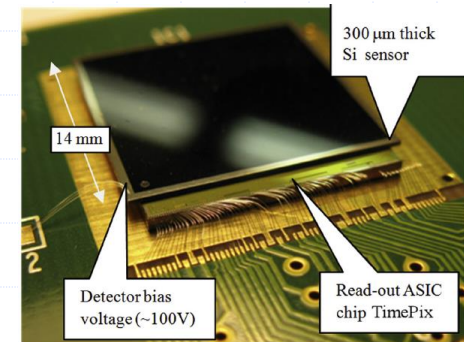
Quantum imaging detection

Active nuclear emulsion

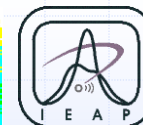
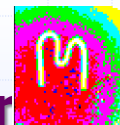
Directional sensitivity



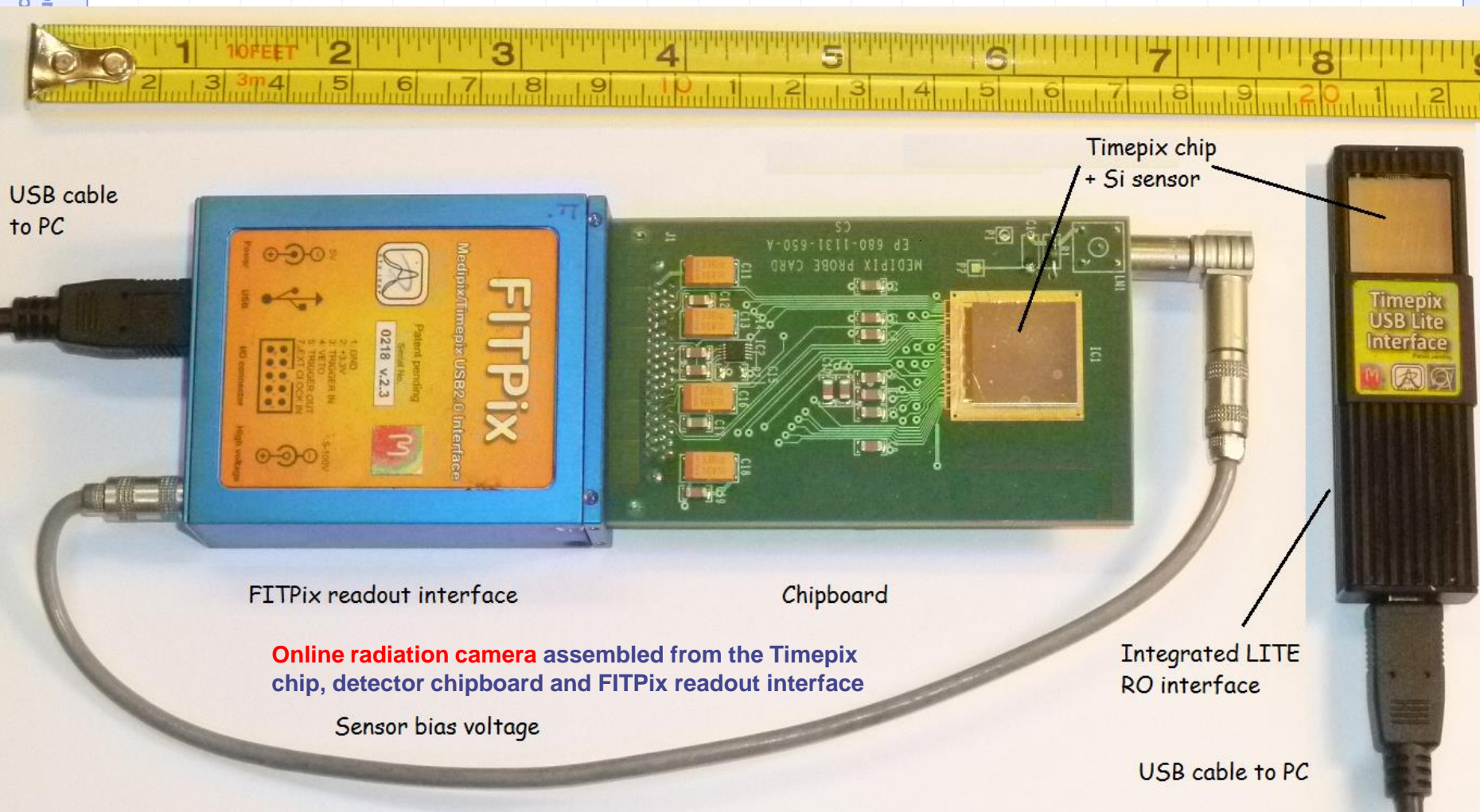
- 256 x 256 pixels (pitch 55 μm)
- 14mm x 14mm = 2 cm^2

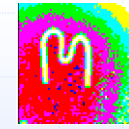


Pixel detectors Medipix/Timepix + Integrated RO electronics + Online & data processing SW + Nuclear Physics know-how: Integrated Radiation Camera



www.cern.ch/medipix

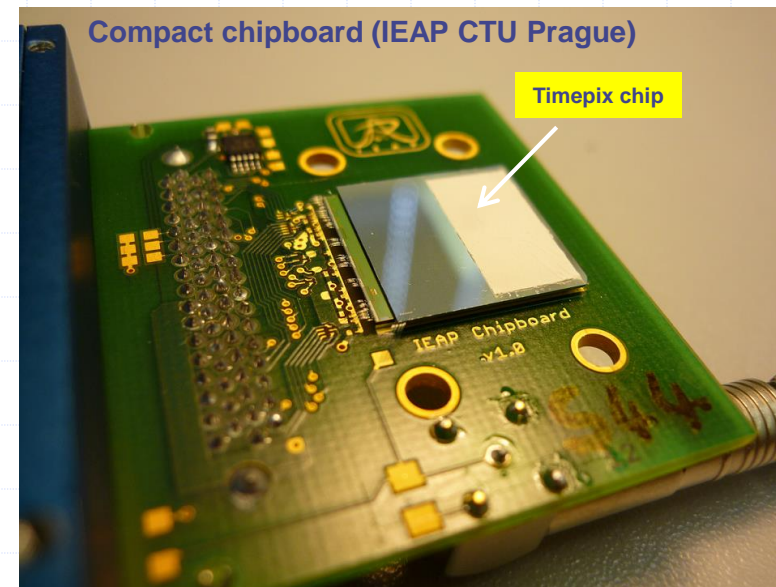
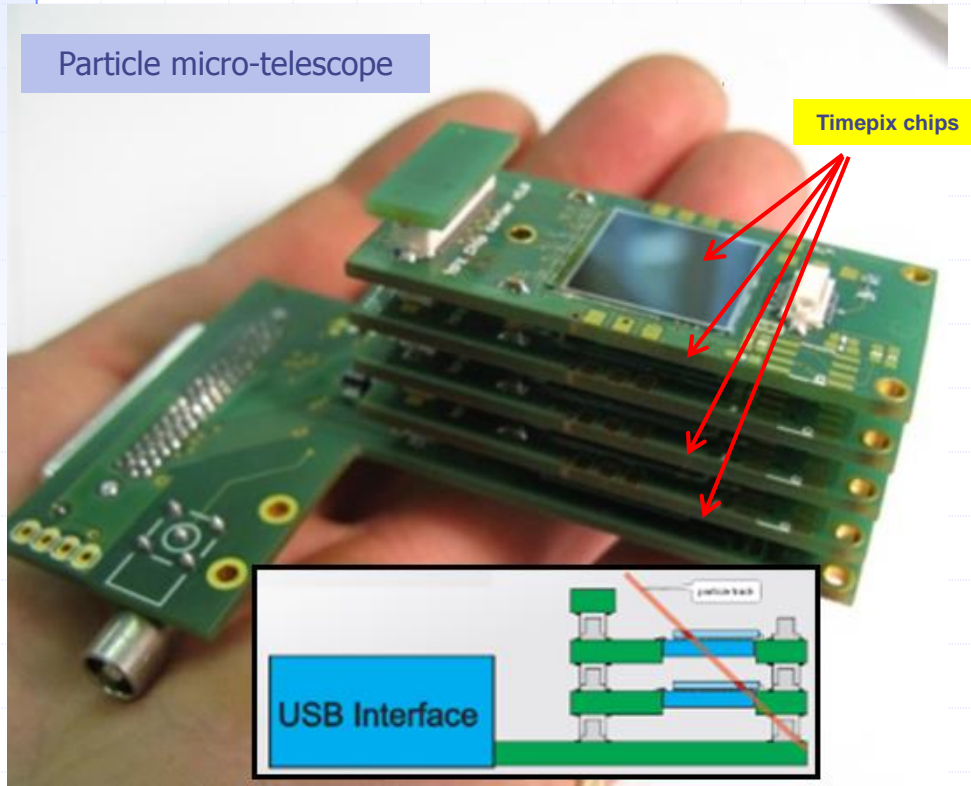




Detector array architectures: Miniaturization, stacking

Enhanced particle tracker (high angular resolution)

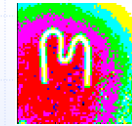
Particle micro-telescope



Particle tracker/telescope assembled/stacked from
four Timepix devices

Detector array architectures:

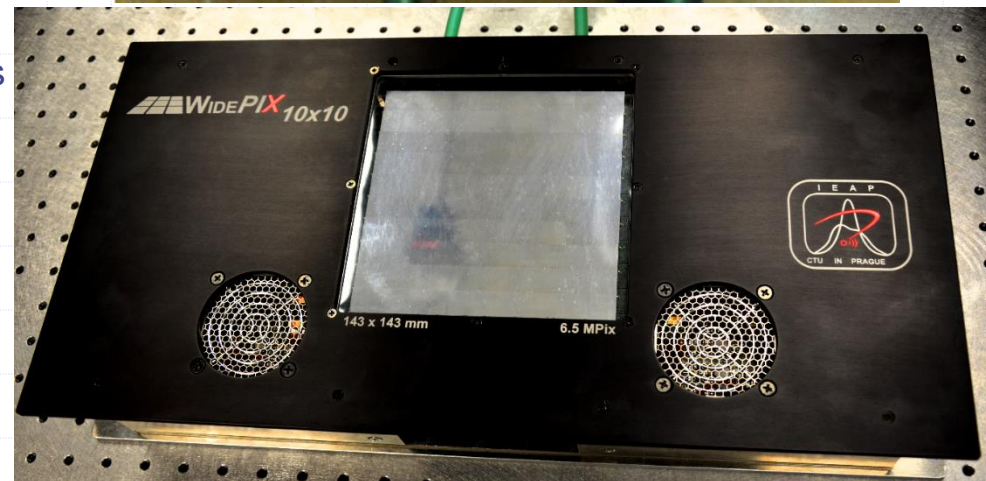
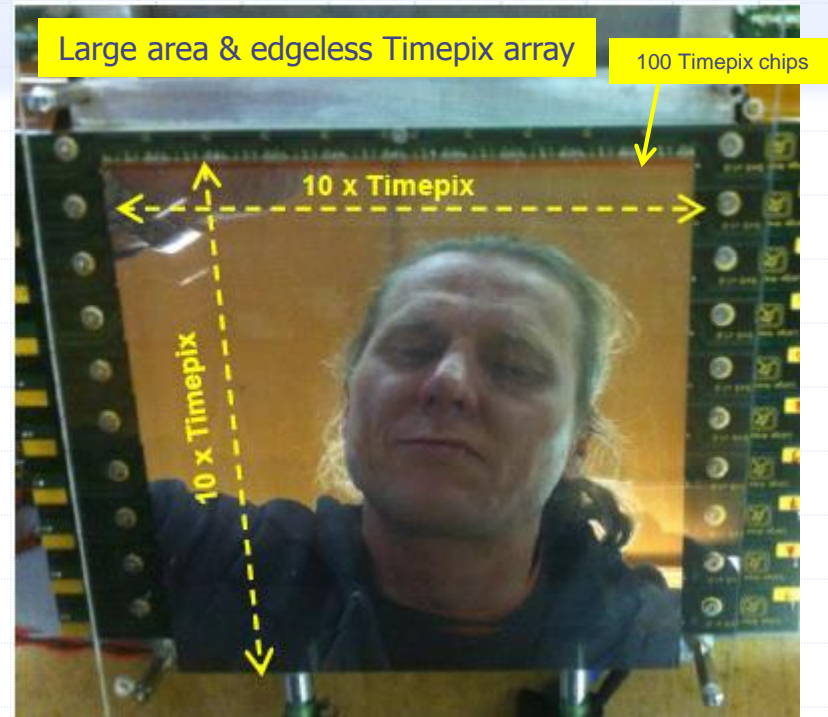
Large sensitive area



- ❑ **WidePIX 10x10 Timepix imager** consists of an array of 100 **edgeless** Timepix detectors (developed in VTT Finland and fabricated by ADVACAM Oy).
- ❑ The whole device was designed, developed and constructed at the IEAP CTU Prague
- ❑ Custom readout electronics + control software tool (Pixelman based)
- ❑ Versions: 10x10, 10x5, 5x4, 10x1, 5x1 chips

Features:

- ❑ Superior image quality without instrumental noise (particle counting)
- ❑ Large (14 cm x 14 cm) **fully sensitive area** with **no gaps** between sensor chips
- ❑ Fully digital detection with **ultra-high contrast** even for light objects (e.g. plastic or soft tissue)
- ❑ Energy discrimination allowing “color” radiography,
- ❑ Compact size and portability (1x PC)
- ❑ Support for major operating systems: Windows, Mac OS, Linux



Detector integrated architectures: Miniaturization + Portability + Remote deployment

RASPIX/Timepix

LITE/Timepix

miniPC, memory, ethernet/WiFi

USB to PC/laptop



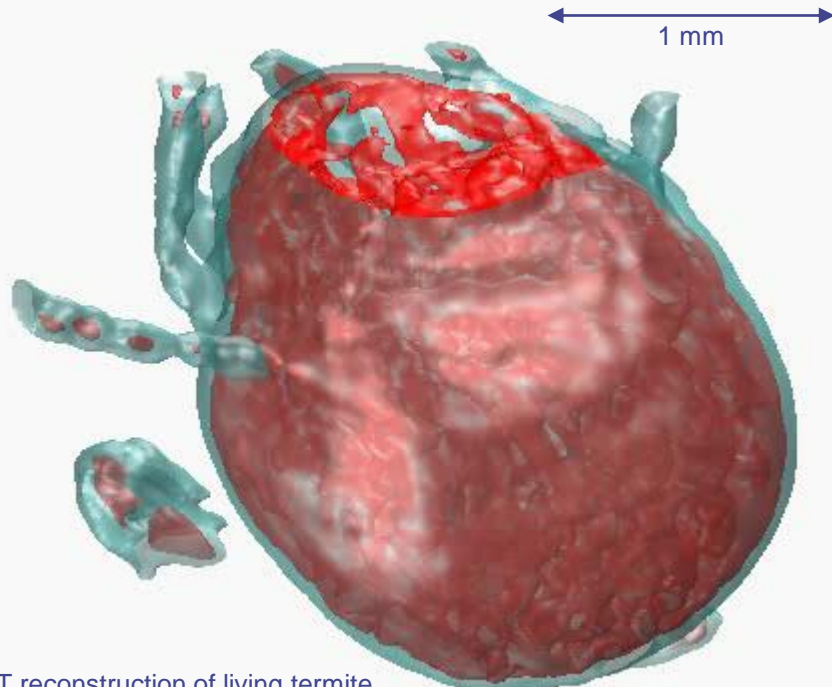
Small animal micro-CT[&] and mixed radiation field characterization in ion beam radiotherapy[#] and in outer space^{*} with pixel detectors Timepix



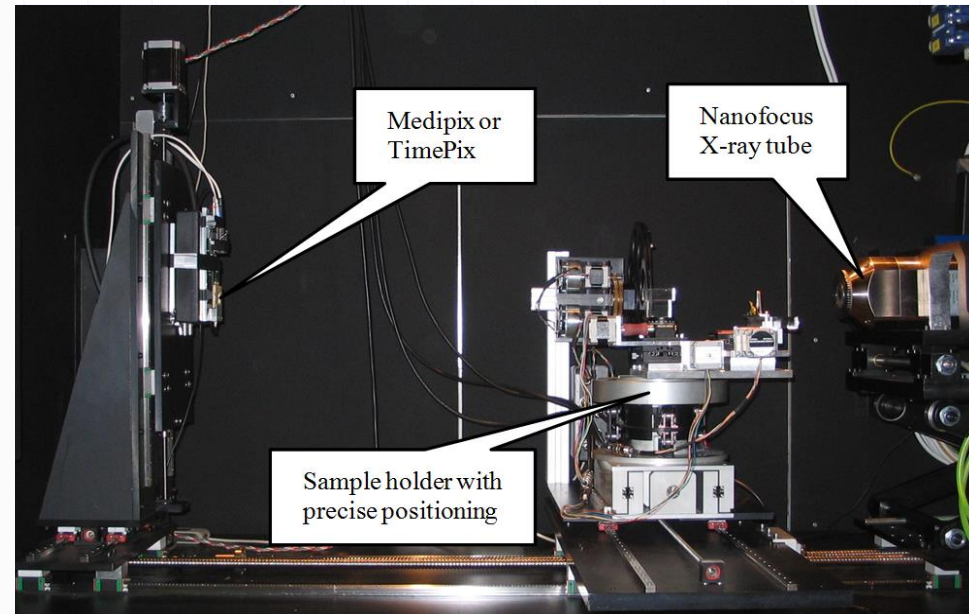
X-ray μ CT: high resolution, soft tissue contrast

In-house laboratory scale system

- ◆ With source-detector distance \rightarrow high magnification + spatial resolution down to 1 μ m.
- ◆ The spatial resolution is currently limited by the X-ray tube focal spot diameter
- ◆ The detector and the X-ray tube are fixed + integrated SW tool
- ◆ Sample placed on the top of a precise multi-axial motorized stage
- ◆ Living specimen + high contrast of soft-tissue samples



CT reconstruction of living termite.
20 projections (5 s each). 300 μ m silicon Timepix detector

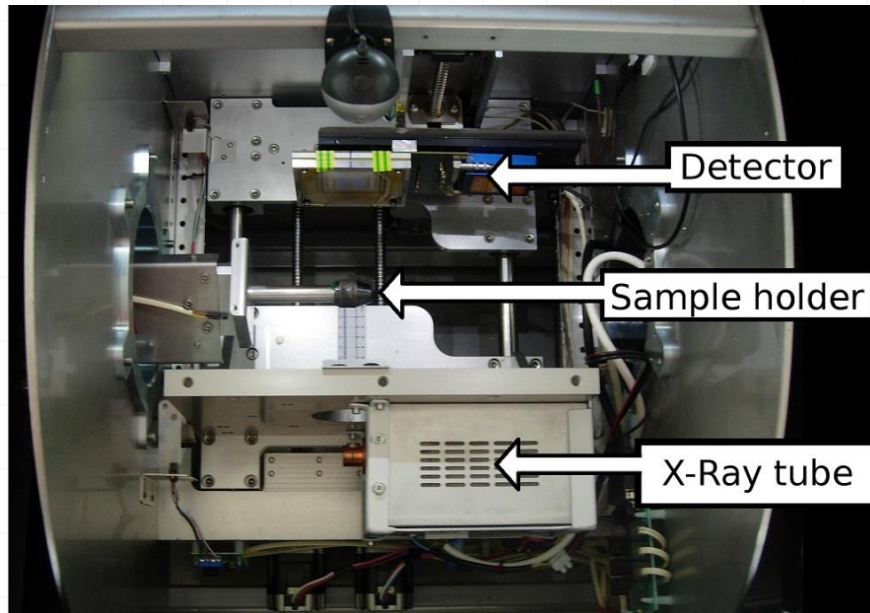




X-ray μ CT: high resolution, soft tissue contrast

Small animal CT scanner

- ◆ Sample fixed at the center of the rotating gantry \rightarrow avoid undesirable movement of biological sample and position uncertainties
- ◆ Built-in house CT system:
 - Timepix Quad detector (512×512 pixels)
 - FITPix interface + processing SW Pixelman
 - KEVEX™ PXS11 X-ray tube μ -focus
 - Custom developed user SW interface
 - Highest achievable spatial resolution $28 \mu\text{m}$



CT scanner rotating gantry: detector, sample holder and radiation source

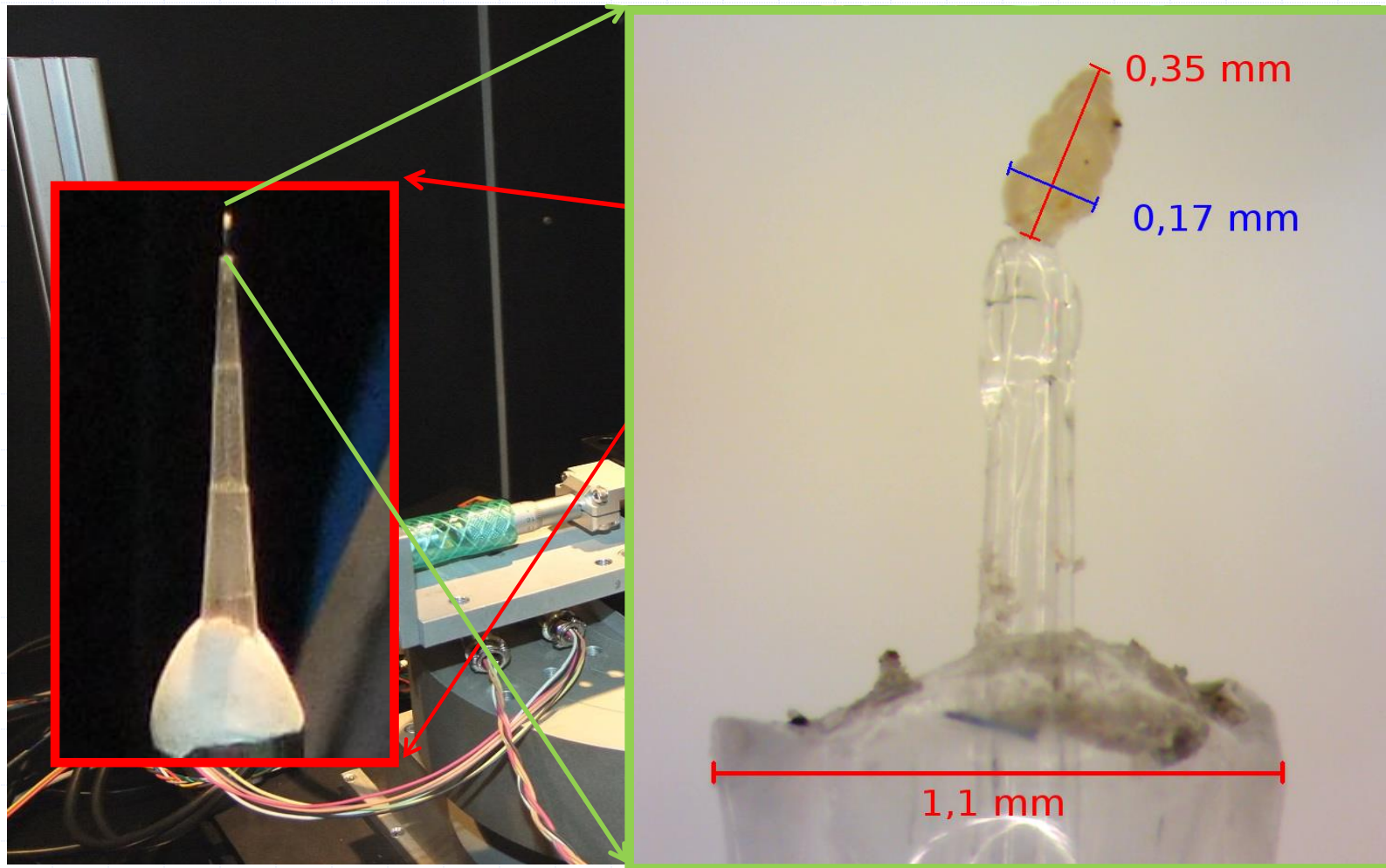


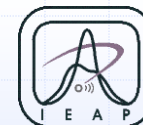
Assembled small animal micro-CT scanner

High resolution high contrast X-ray μ CT

Organic samples with micro-structures

- ◆ Radiography of sea microfossils (foraminifera)

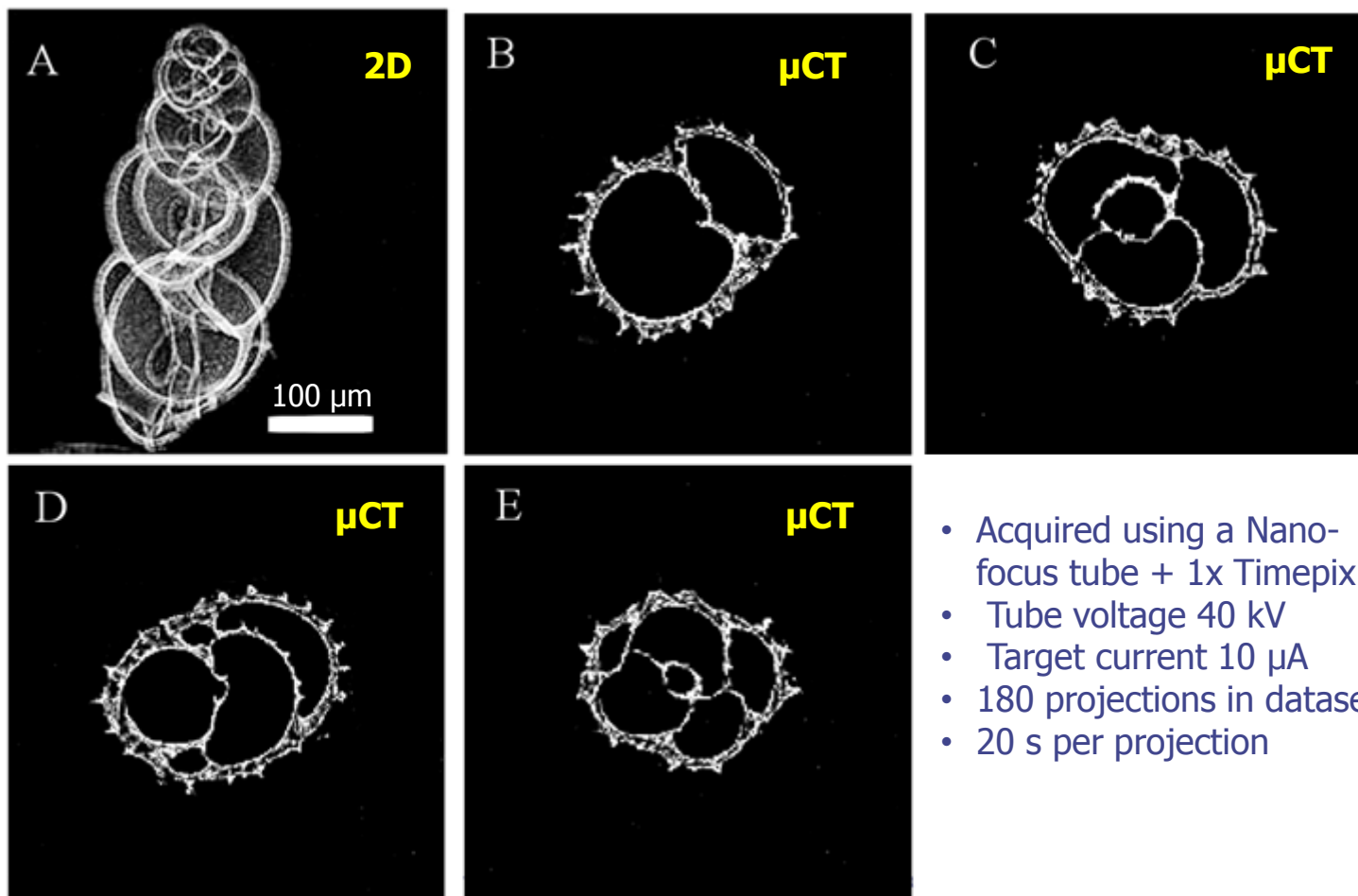




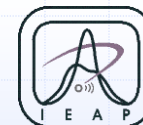
High resolution high contrast X-ray μ CT

Organic samples with micro-structures

- ◆ A) Example of radiographic projection; B – E) Virtual sections through the reconstructed voxel model



- Acquired using a Nano-focus tube + 1x Timepix
- Tube voltage 40 kV
- Target current 10 μ A
- 180 projections in dataset
- 20 s per projection

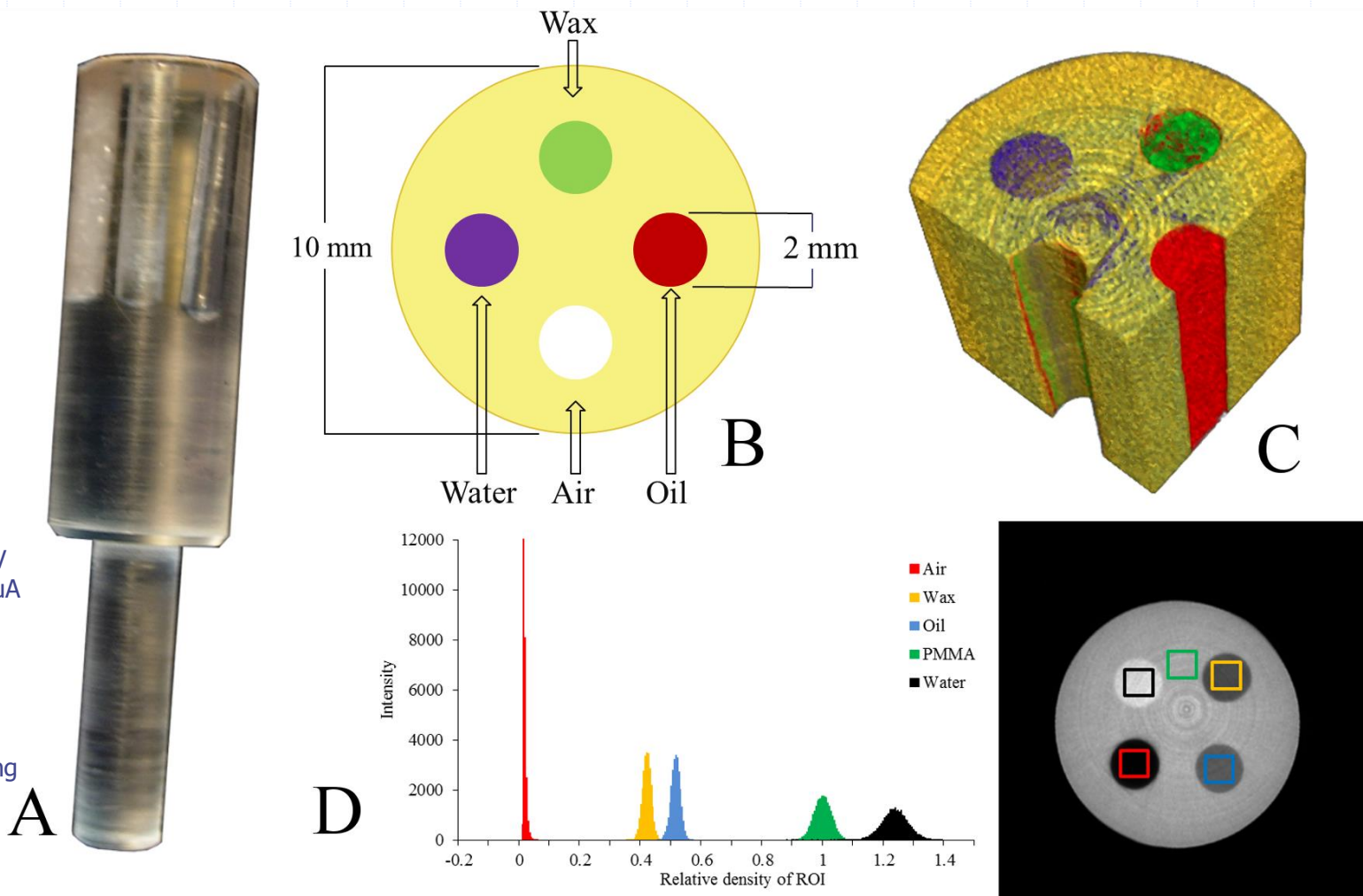


High resolution high contrast X-ray μ CT

Soft tissue phantom + density resolving power

- Small animal uCT
- QUAD Timepix
- Tube voltage 60 kV
- Tube current 100 μ A
- 180 projections
- 20 s per projection

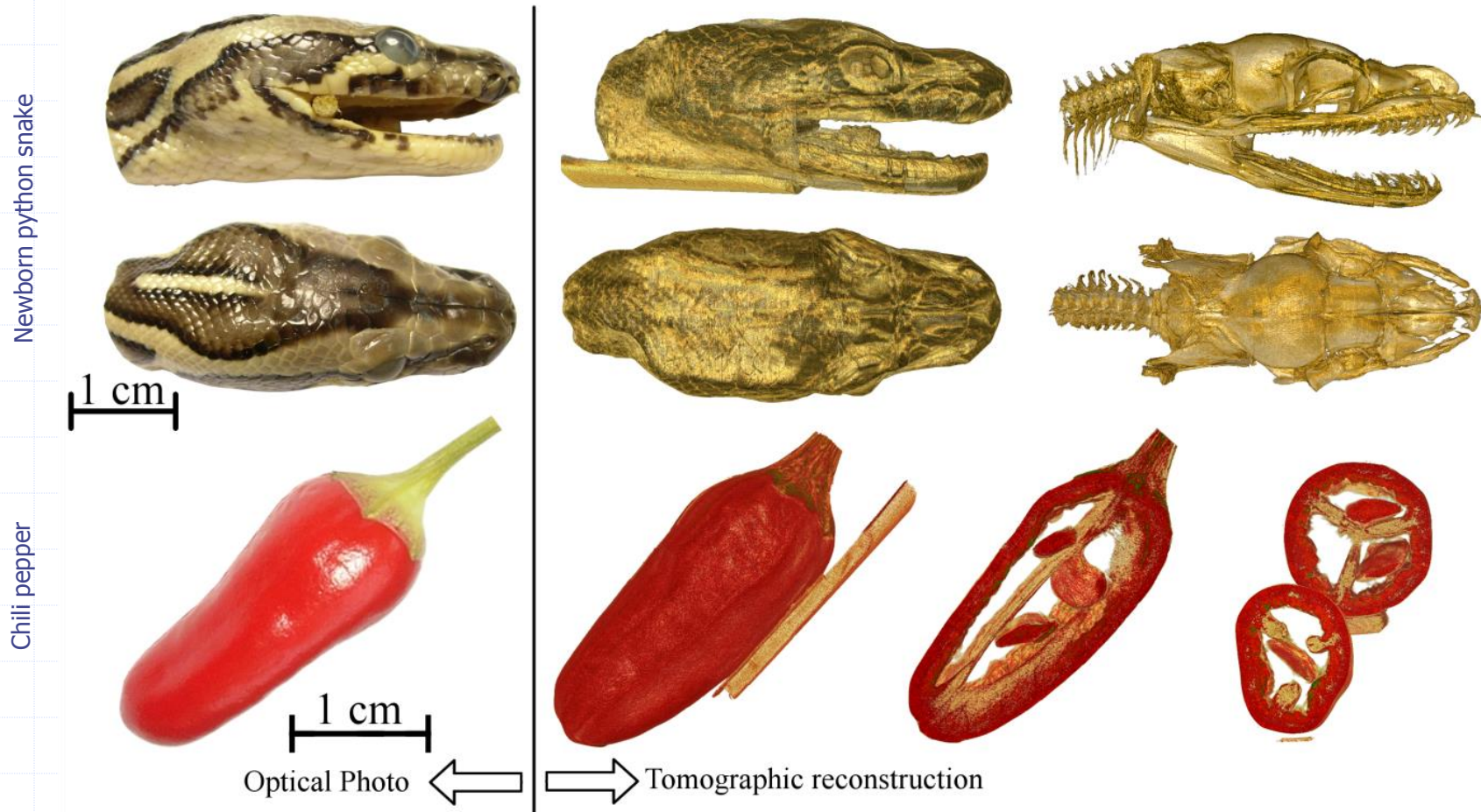
- A: optical photo
- B: scheme layout
- C: volume rendering
- D: relative density histogram



μ CT slice

High resolution high contrast X-ray μ CT

Soft tissue imaging of complex biological objects



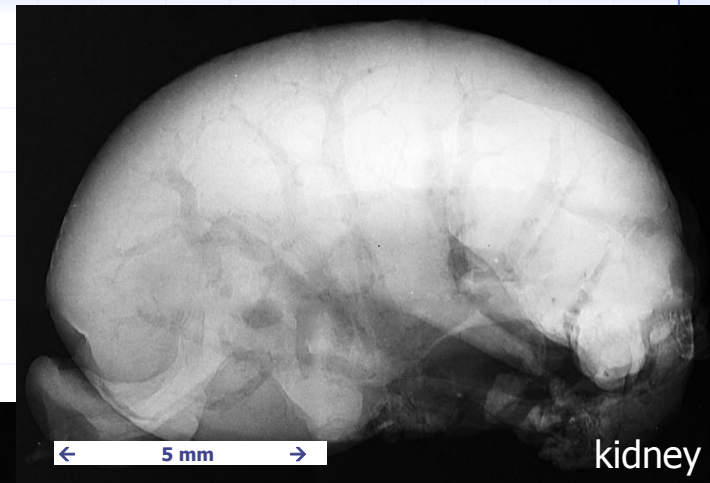
- Small animal μ CT + Timepix QUAD (512 x 512 pixels), silicon 300 μ m sensor
- X-ray tube voltage 60 kV, current 90 μ A, 180 projections, 10 s per projection.



High resolution high contrast X-ray imaging

Soft tissue imaging of complex biological objects

- ◆ 2D X-ray imaging of native and alcohol-fixed rat and mouse inner organs with the aim to study their anatomy and morphology
- ◆ Examples of alcohol-fixed mouse organs without any contrast agents



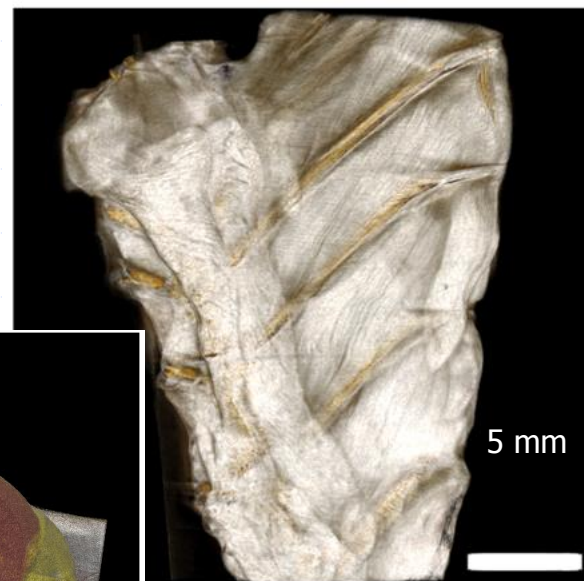
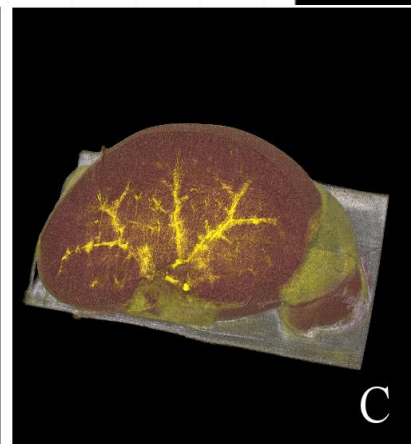
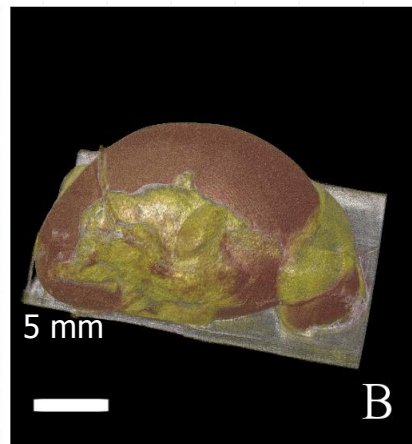
- Samples shown scanned with the small animal μ CT scanner
- Timepix Quad detector
- Spatial resolution 28 μ m
- Acquisition parameters: Tube voltage 60kV, current 100 μ A, 45 s/acq

High resolution high contrast X-ray imaging

Imaging of biological objects + contrast agents

- ◆ Different available contrast staining methods and commercially available in vivo dedicated contrast agents are tested
 - Iodine based staining
 - BaSO₄
 - Aurovist™
- ◆ small animal μ CT scanner + Timepix Quad detector

Volume rendering of a rat kidney with contrast enhanced by Aurovist™:
 A) Optical photo of the sample; B) Volume rendering of the tomographic reconstruction; C) A virtual section through rendered volume revealing enhanced contrast enhancement of cavities.

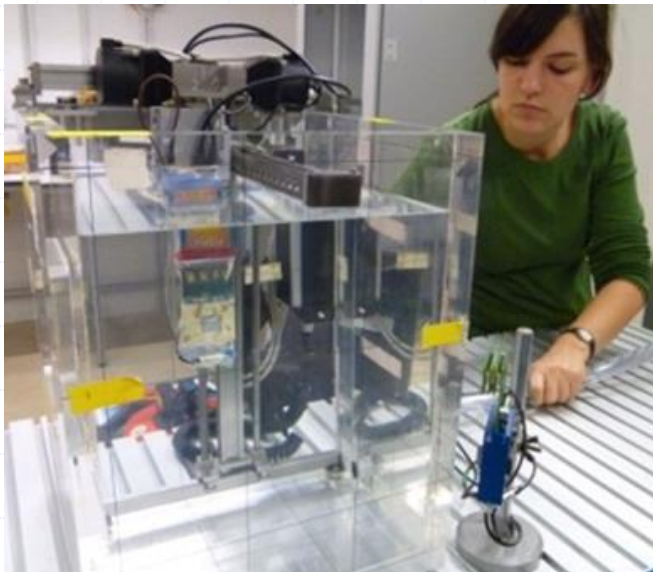


Volume rendering of a part of laboratory rat chest stained with Lugol's solution reveals the fibrous muscle structure.

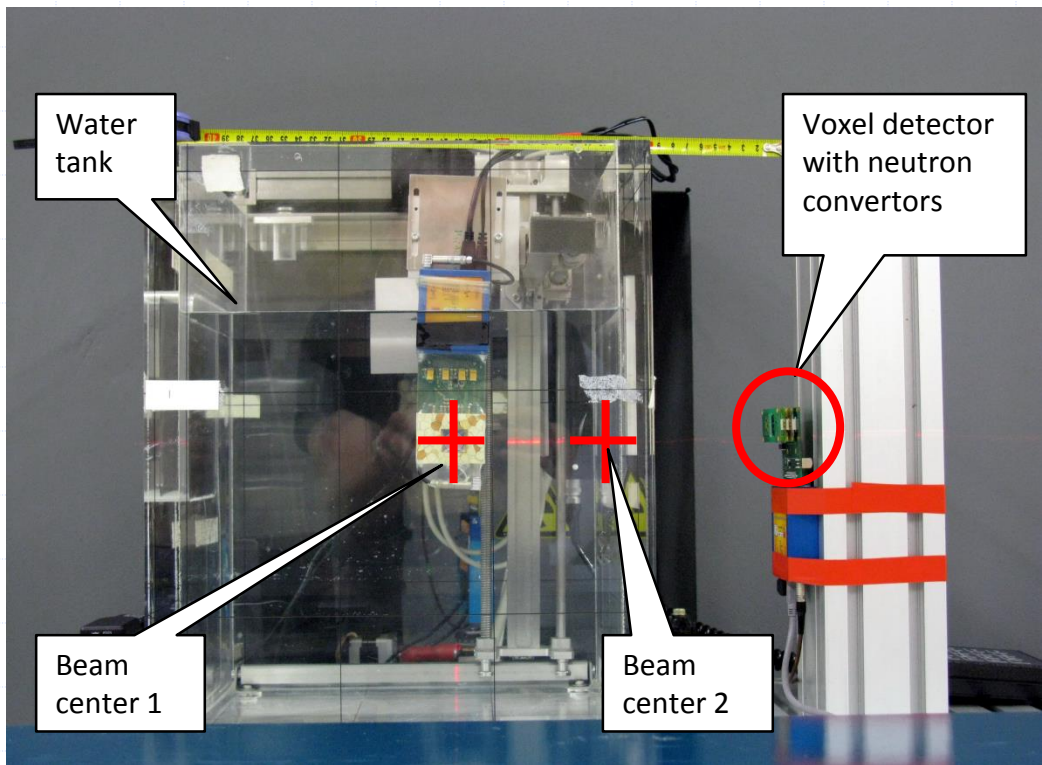
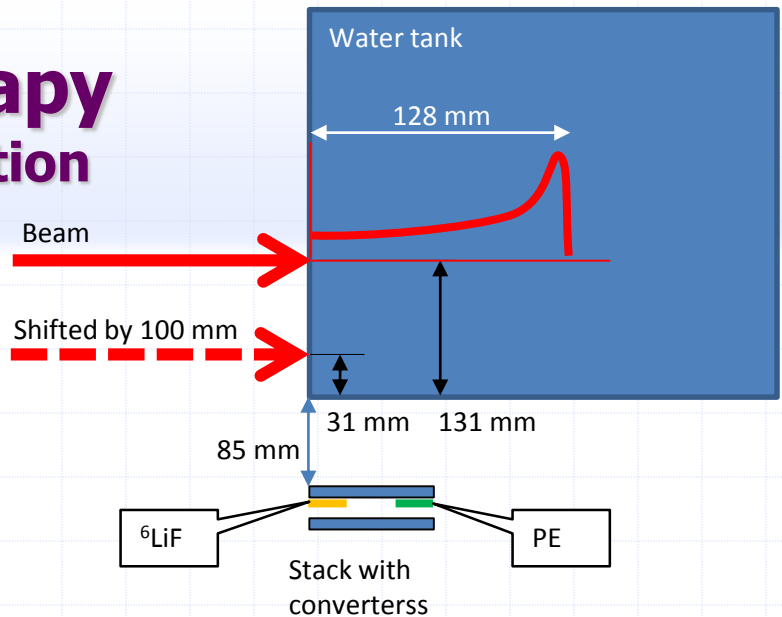
Small animal micro-CT& **mixed radiation field characterization in ion beam radiotherapy[#] and in outer space^{*} with pixel detectors Timepix**

Ion beam radiotherapy

Imaging of secondary radiation



Beams: 1H, 4He, 12O, 16O
E: 50-400 MeV/u



Ion beam radiotherapy

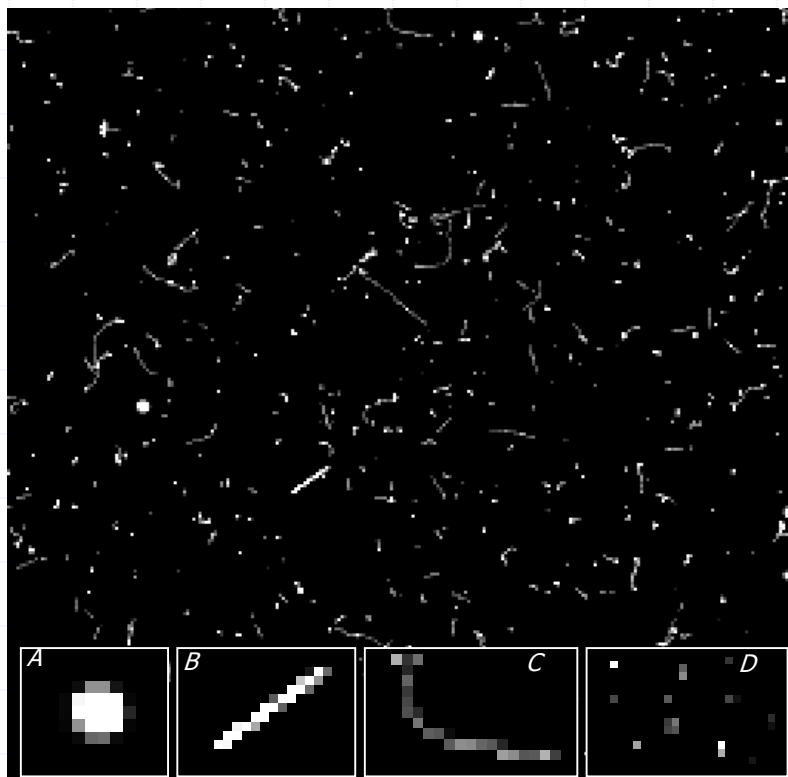


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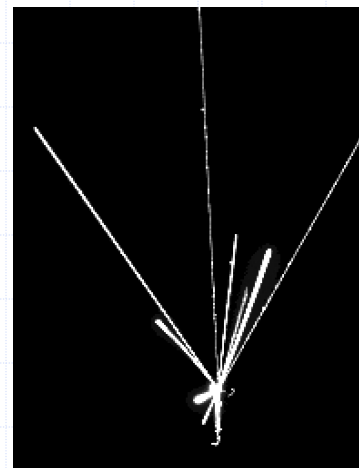
GERMAN
CANCER RESEARCH CENTER
IN THE HELMHOLTZ ASSOCIATION



Particle tracking with pixel detectors Timepix



Lab radioactive source + cosmic ray background
+ long exposure



HIT: 430 MeV/u 12C
primary beam: high LET
interaction

HIT: 430 MeV/u 12C
primary beam:
energetic/delta electrons



Proton track

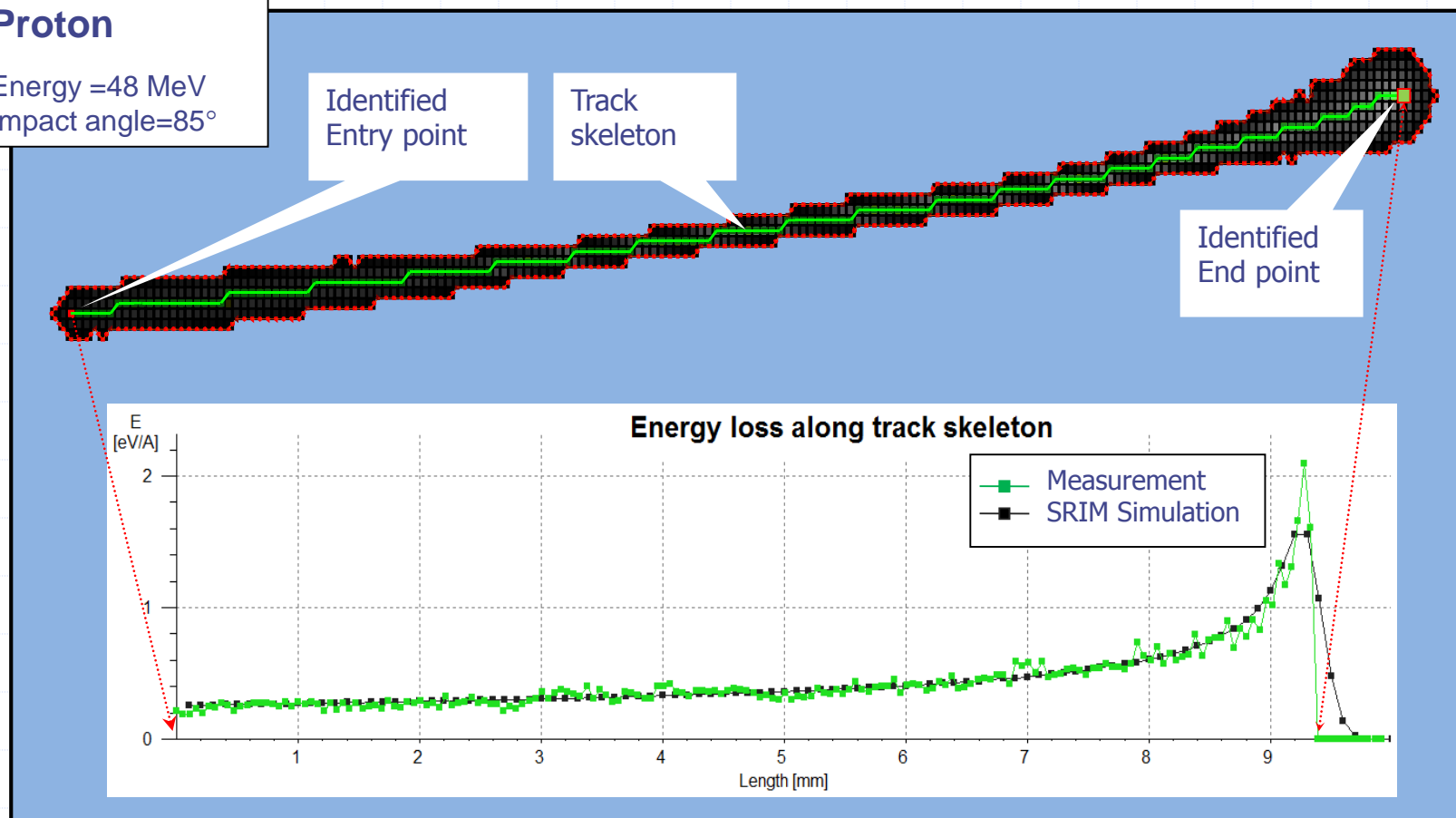
HIT: 48 MeV proton primary beam

Proton

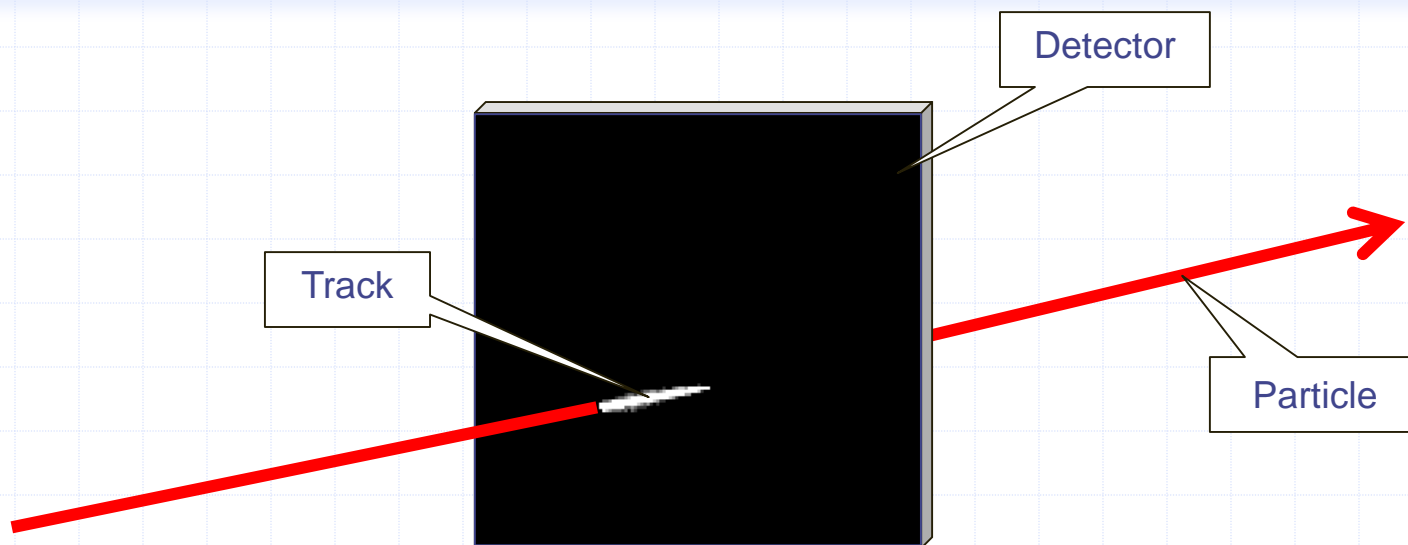
 Energy = 48 MeV
 Impact angle = 85°

 Identified
 Entry point

 Track
 skeleton

 Identified
 End point


Calculation LET



- ◆ LET calculated as the ratio of total deposited energy (integral of energies in all pixels of the cluster) and path length across of the particle through the sensor volume.

$$LET = \frac{dE}{dx}$$

- ◆ This calculation is affected of the path length which is better for long clusters.

Ion beam radiotherapy

Imaging + setup of measurements



Exit window

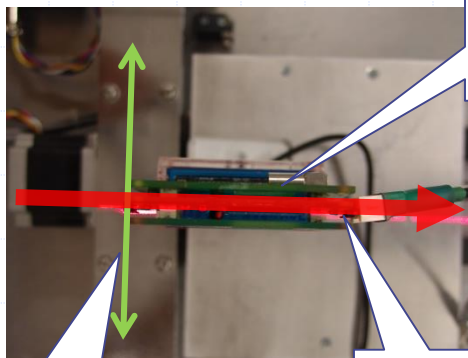
Detector
Timepix



- ◆ Parameters of beam
 - Type : ^{12}C
 - Energy: 430 MeV/u
(5120 MeV)
 - FWHM: 9.8 mm

Beam

View from the top



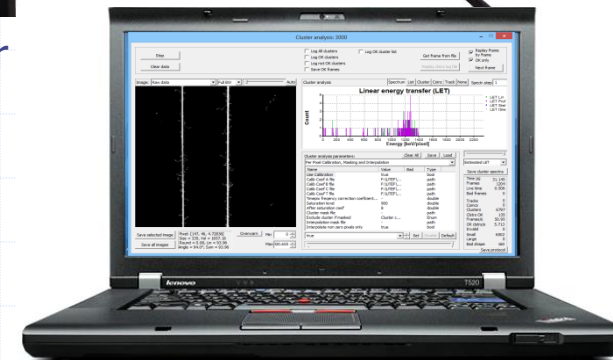
Detector
Timepix

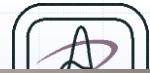
Position
system

- ◆ Parameters of detector
 - Silicon detector
 - 1000 μm thickness
 - BIAS: 450 V
 - TOT mode
 - Event by event

Position
system

Beam

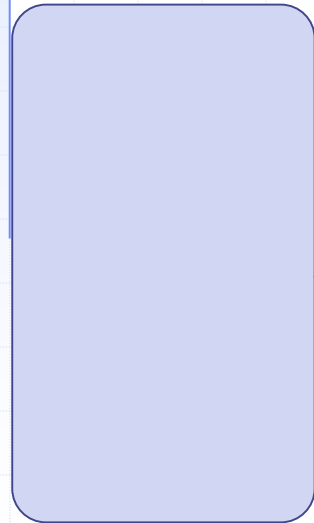
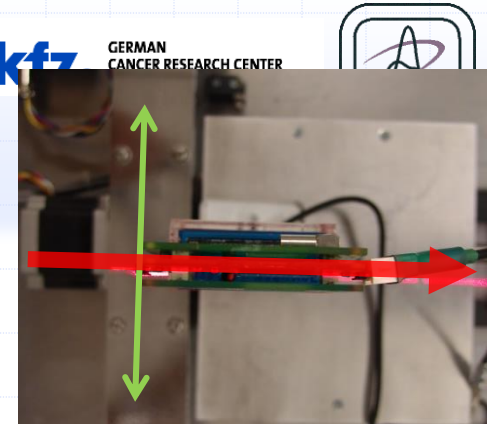




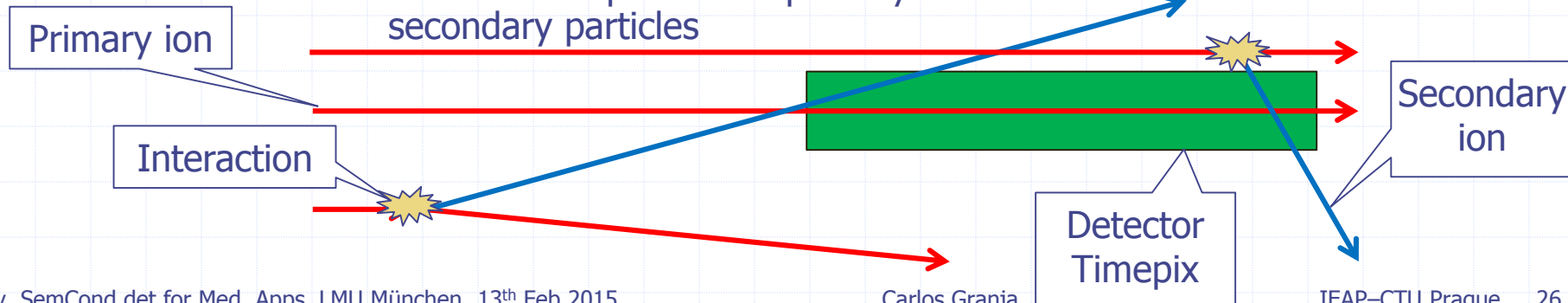
Setup of measurement

View from the top

- ◆ The detector orientation was parallel with respect to the beam axis.
- ◆ 8 different positions

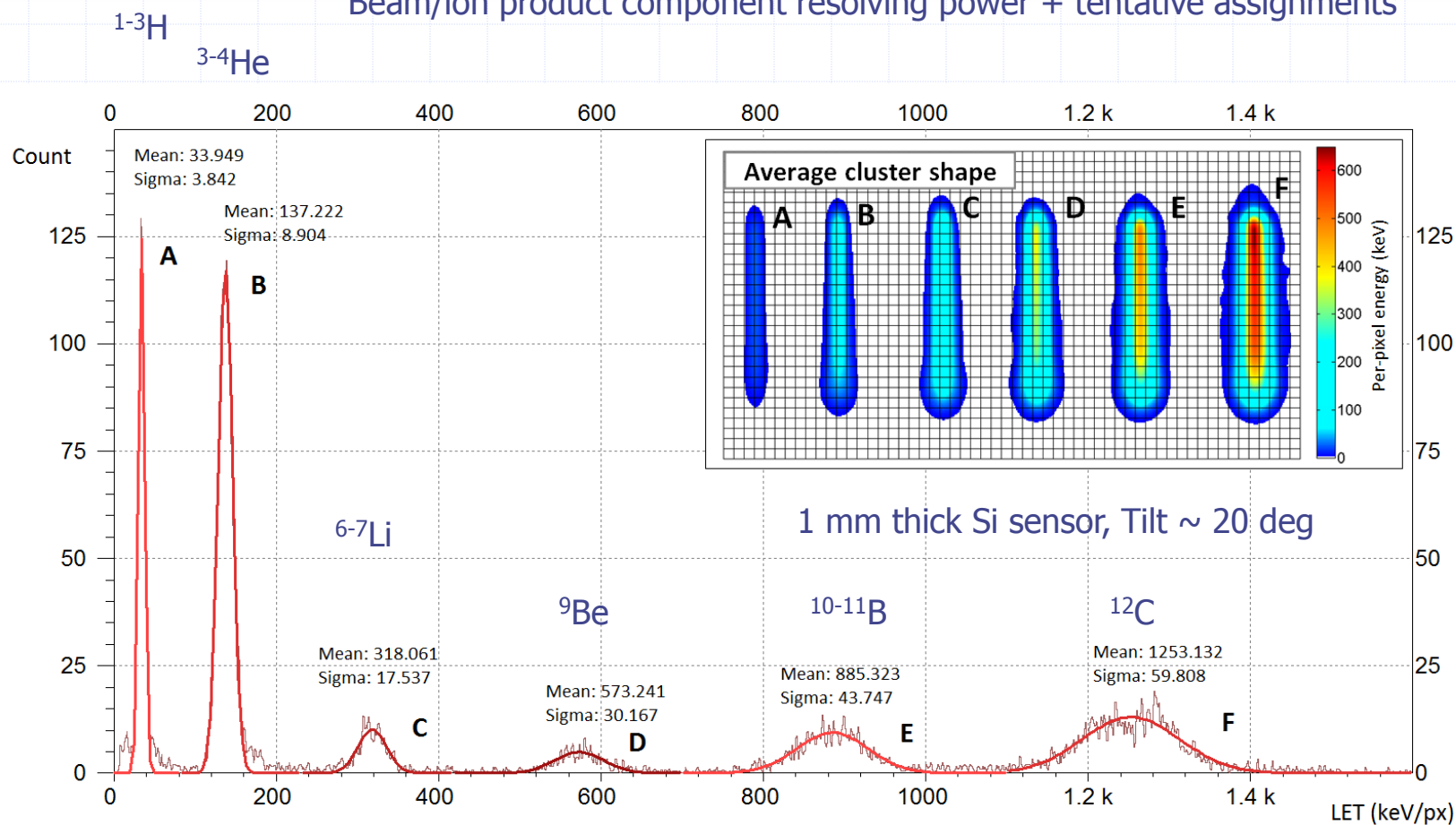


- ◆ Detection of the primary ions
- ◆ Orientation of the detector was chosen in order to separate the primary and secondary particles



Ion groups sensitivity LET spectrum

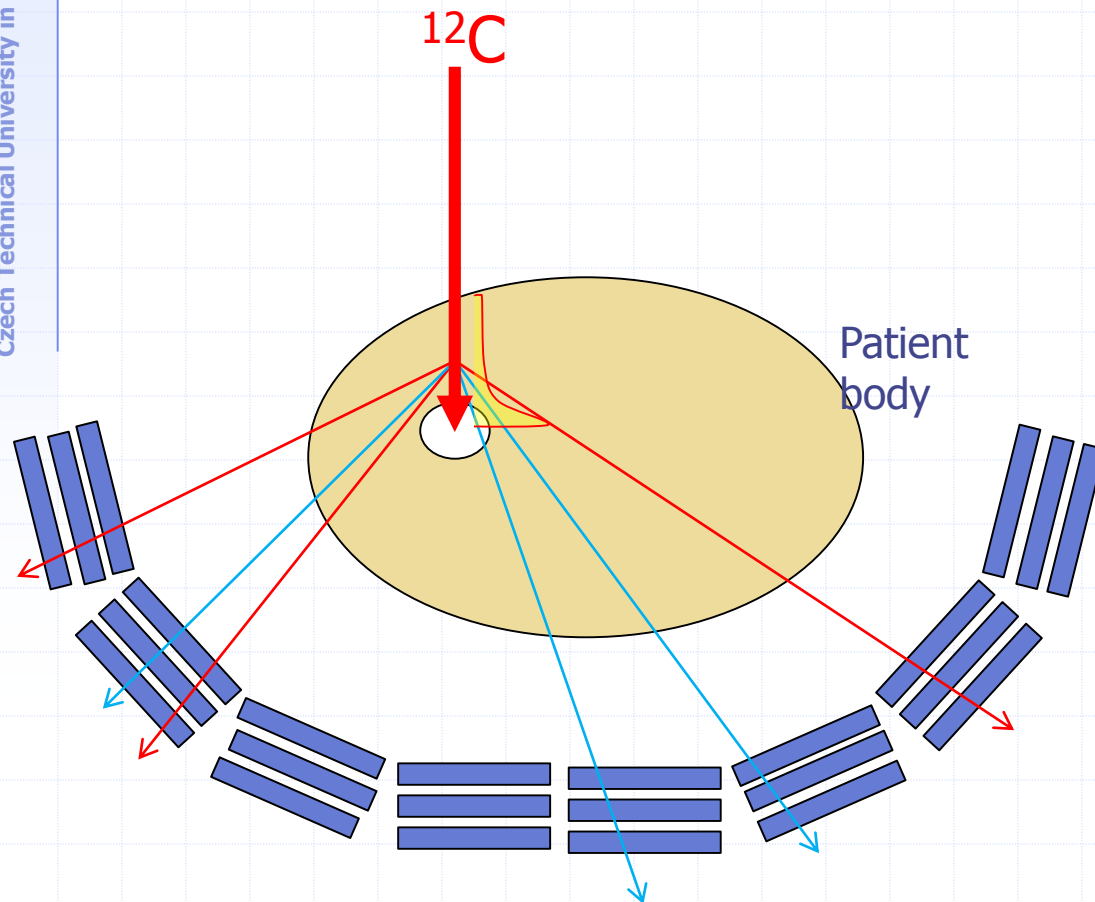
Beam/ion product component resolving power + tentative assignments



LET spectrum of particles detected 20 mm from the beam center.

Imaging for hadron therapy

Tracking of secondary particles & vertex visualization

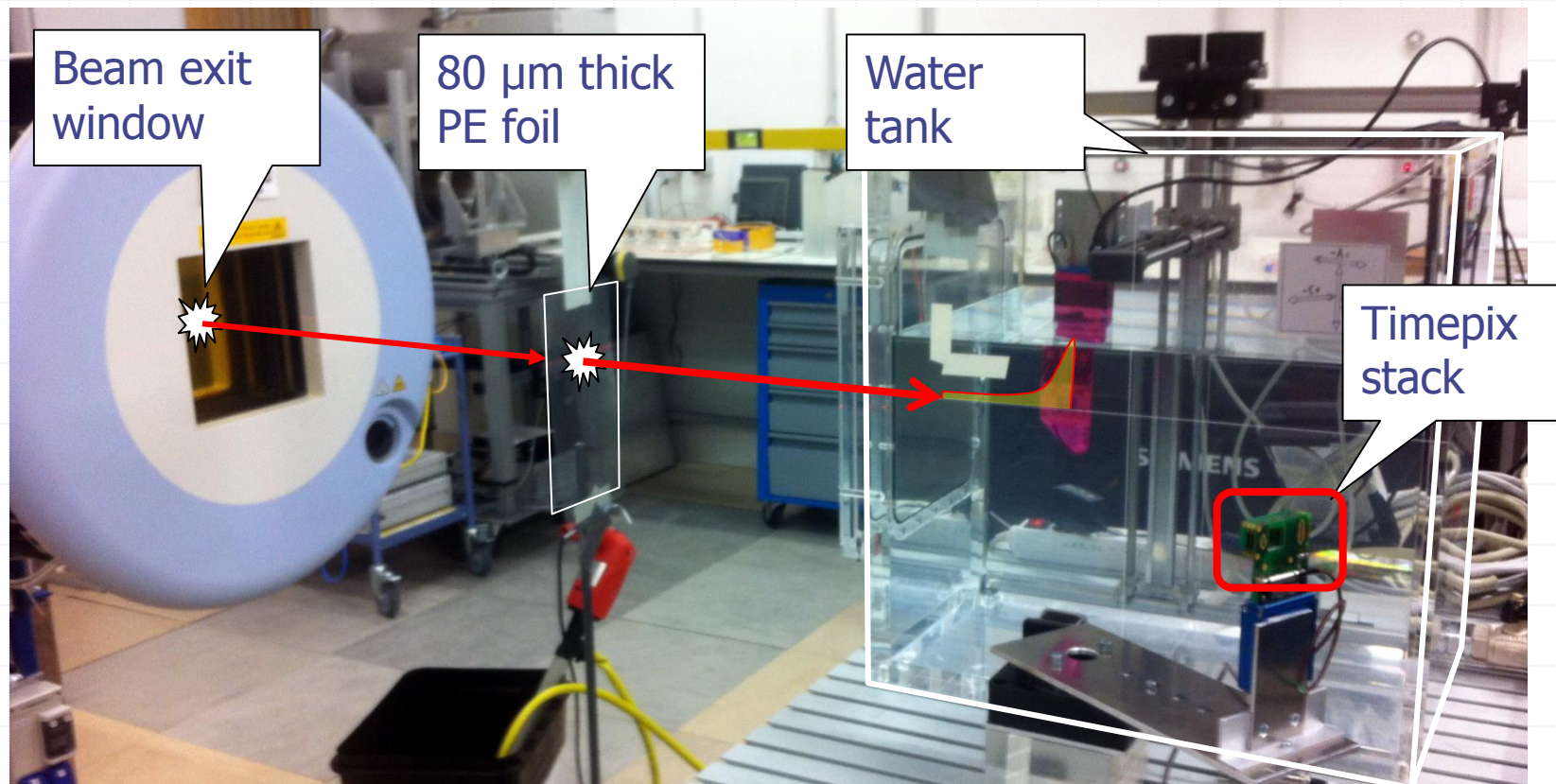


- ◆ The tracker can be scaled to surround the irradiated object.
- ◆ Tracker data can be back-projected to form an image of the beam path.
- ◆ Possibility to select particles with higher penetration power would improve quality.

Imaging for hadron therapy

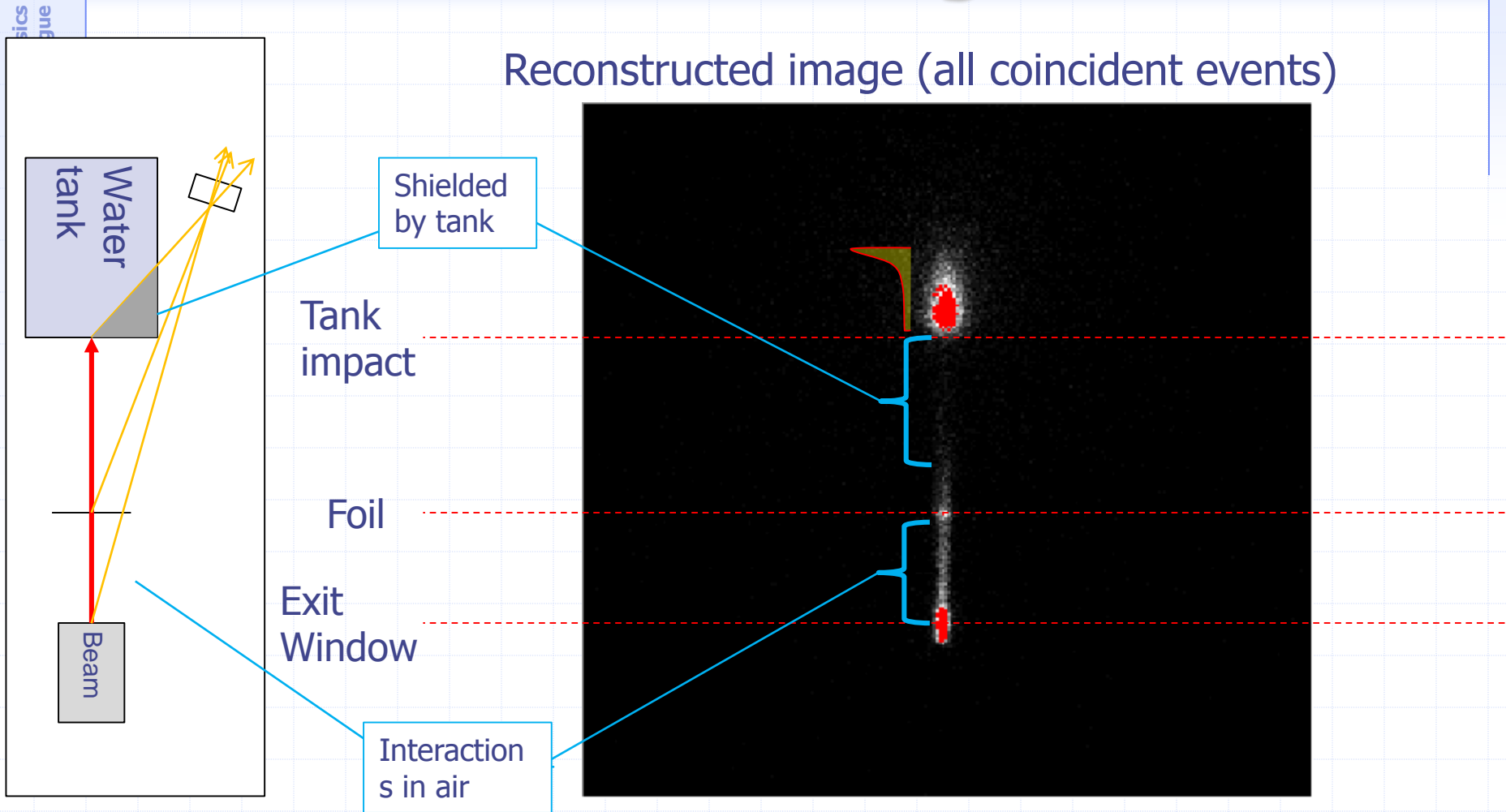
Tracking of secondary particles & vertex visualization

Timepix stack (voxel) detector + experimental setup



Vertex visualization

Beam line can be imaged



Small animal micro-CT& **mixed radiation field characterization in ion beam radiotherapy[#] and in outer space^{*} with pixel detectors Timepix**

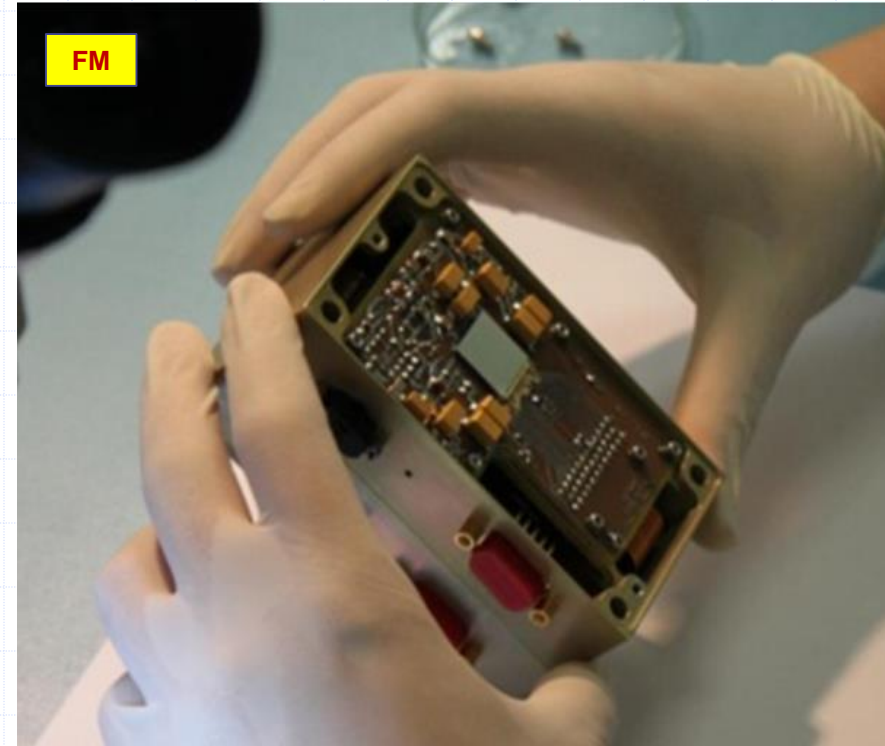
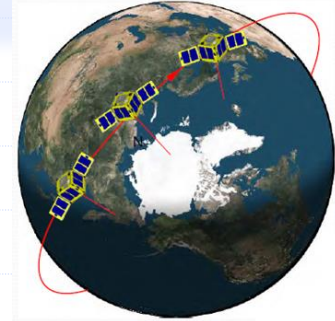
Spacecraft Payload SATRAM

Space Application of Timepix Radiation Monitor



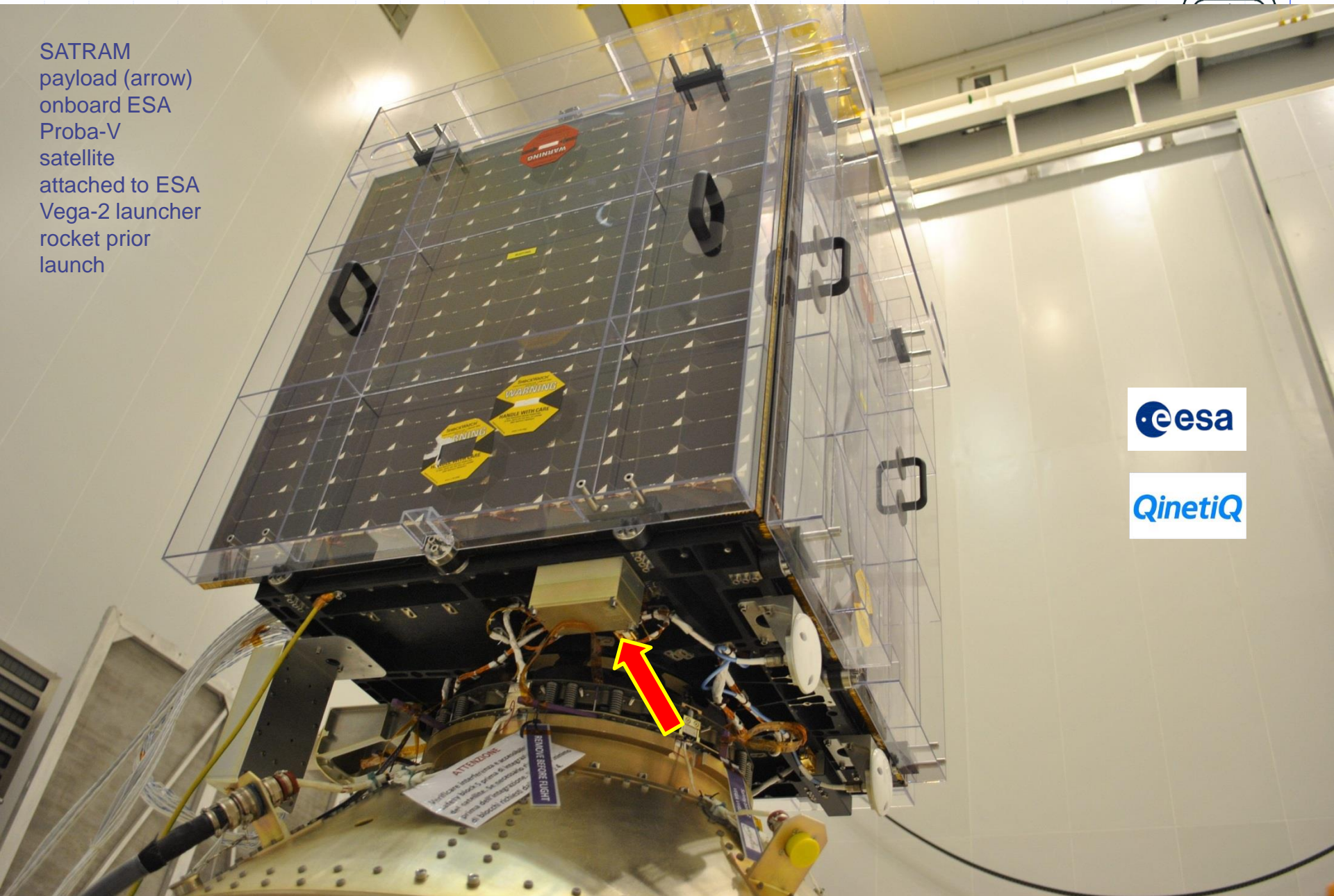
Characterization of space radiation in Low Earth Orbit (LEO) onboard ESA PROBA-V satellite

- ◆ Altitude ~ 820 km, polar sun synchronous orbit, 82° inclination
- ◆ Timepix for the first time in **open space** – currently TRL 9
- ◆ Launched 7th May 2013



Size: 10.8 cm × 6.3 cm × 5.6 cm, full mass 340 g

SATRAM
payload (arrow)
onboard ESA
Proba-V
satellite
attached to ESA
Vega-2 launcher
rocket prior
launch





ESA Vega-2 rocket



- ❑ SATRAM payload (arrow) onboard ESA Proba-V satellite.
- ❑ ESA Vega-2 launcher rocket upper stage



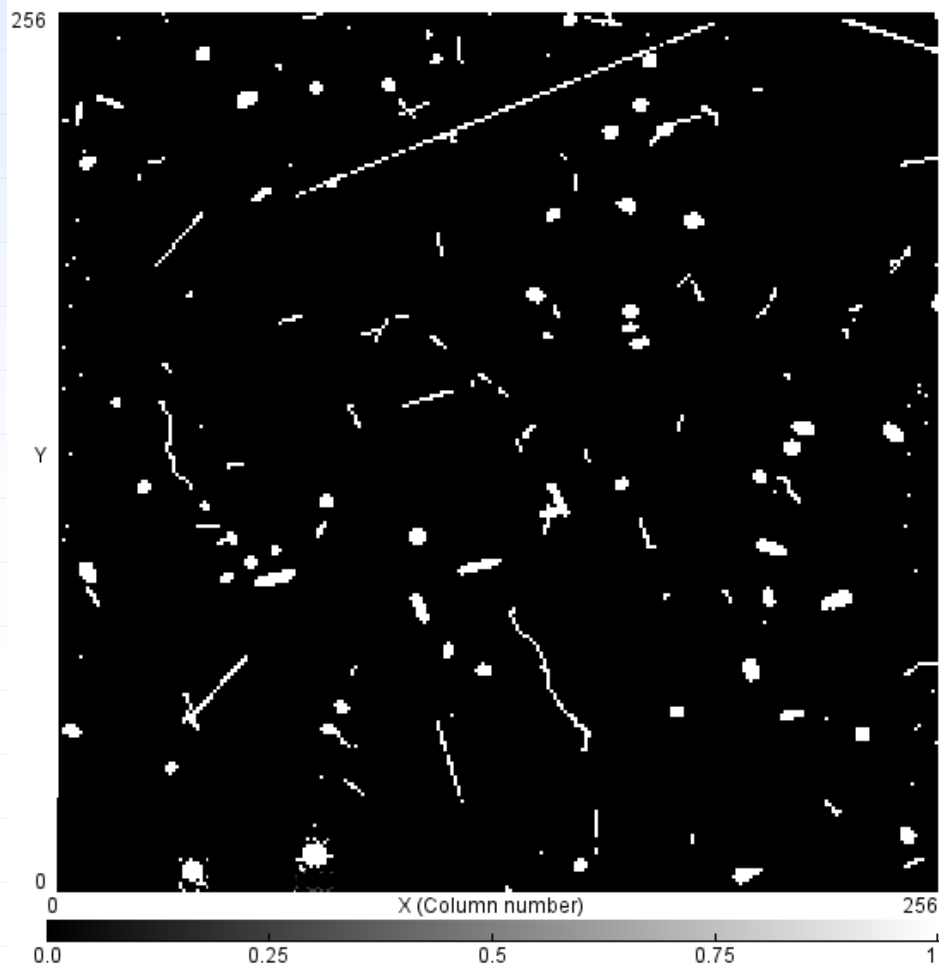
Timepix SATRAM/ESA Proba-V in Open Space

Quantum imaging detection/monitoring of space radiation

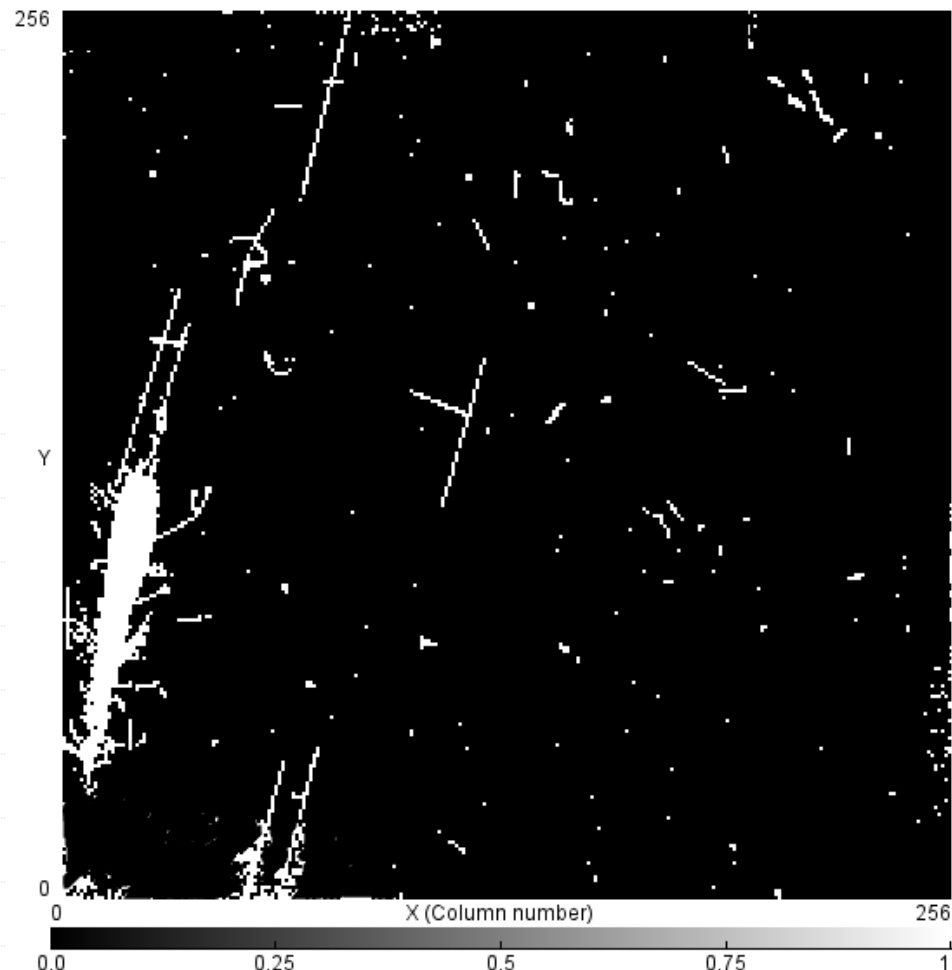
physics
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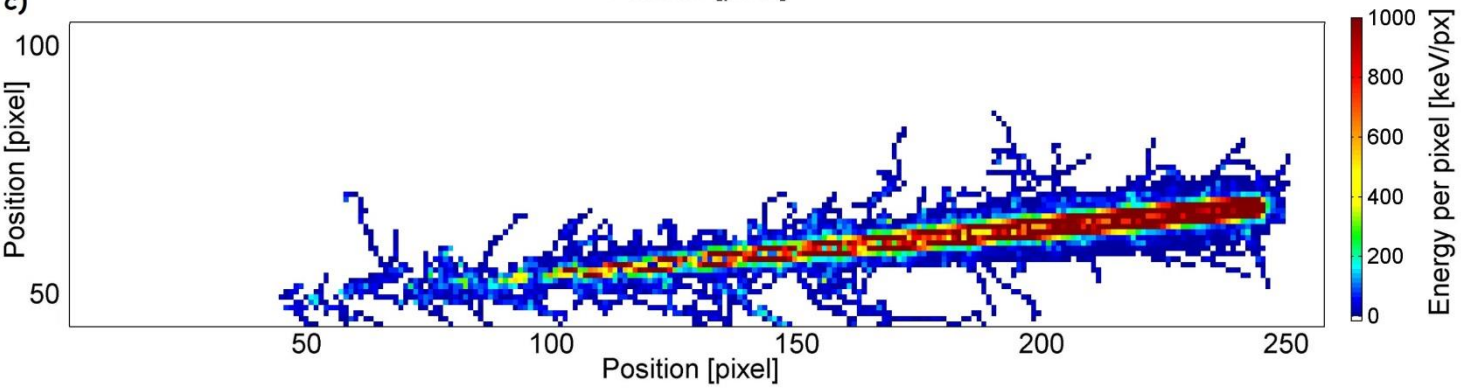
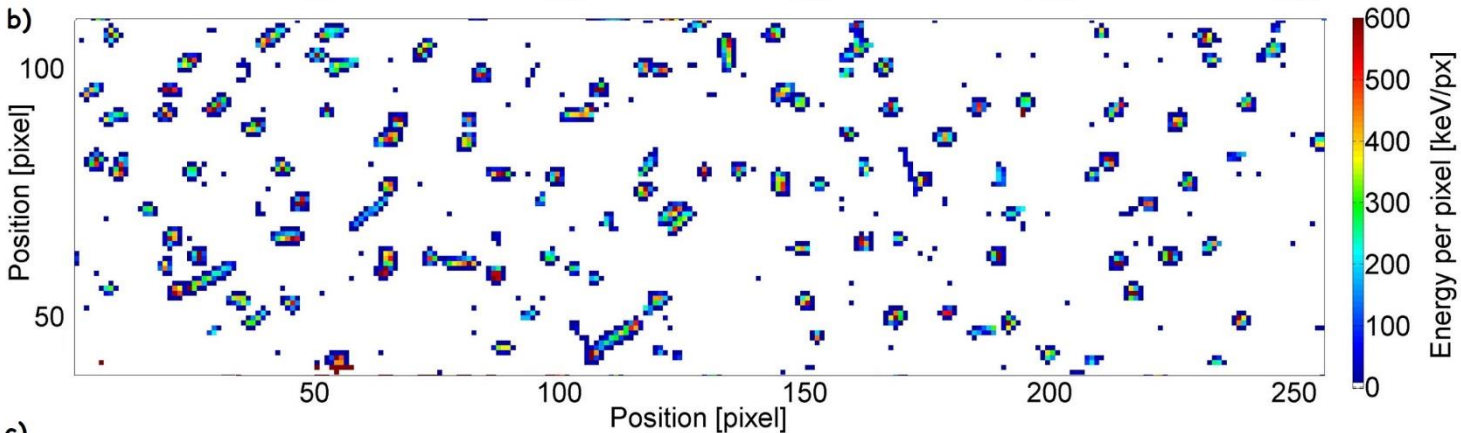
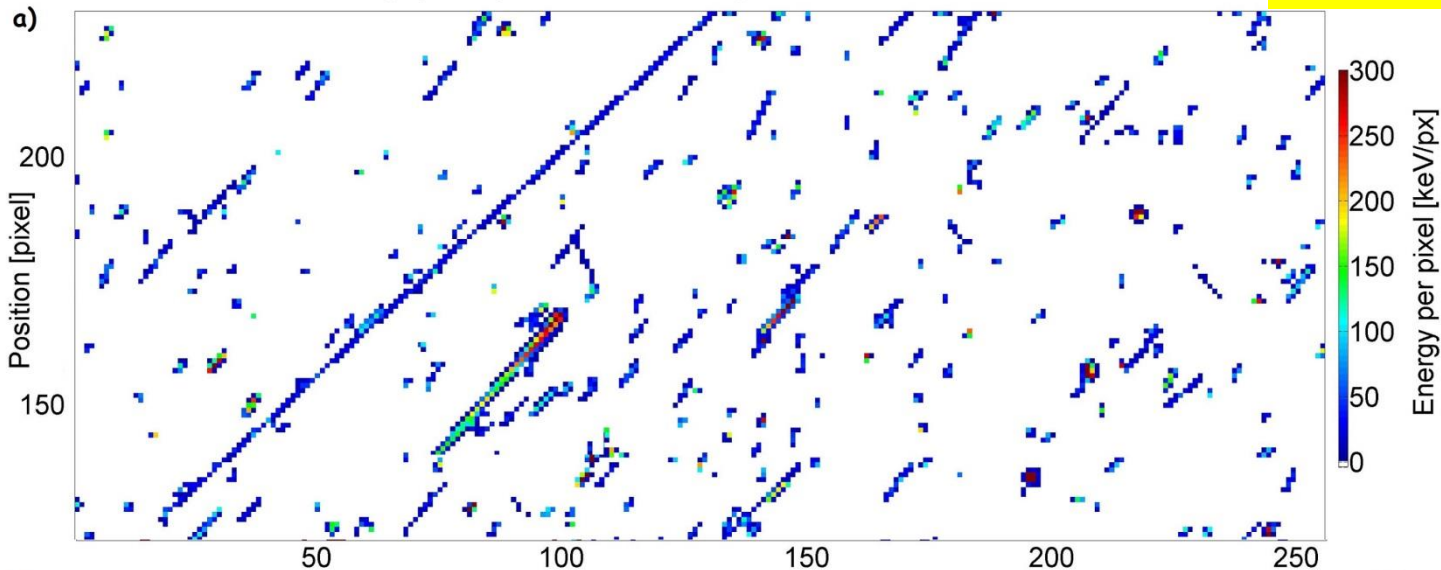
Low Earth Orbit (LEO), 820 km altitude

11.11.2013 12:39:17



11.11.2013 11:16:59





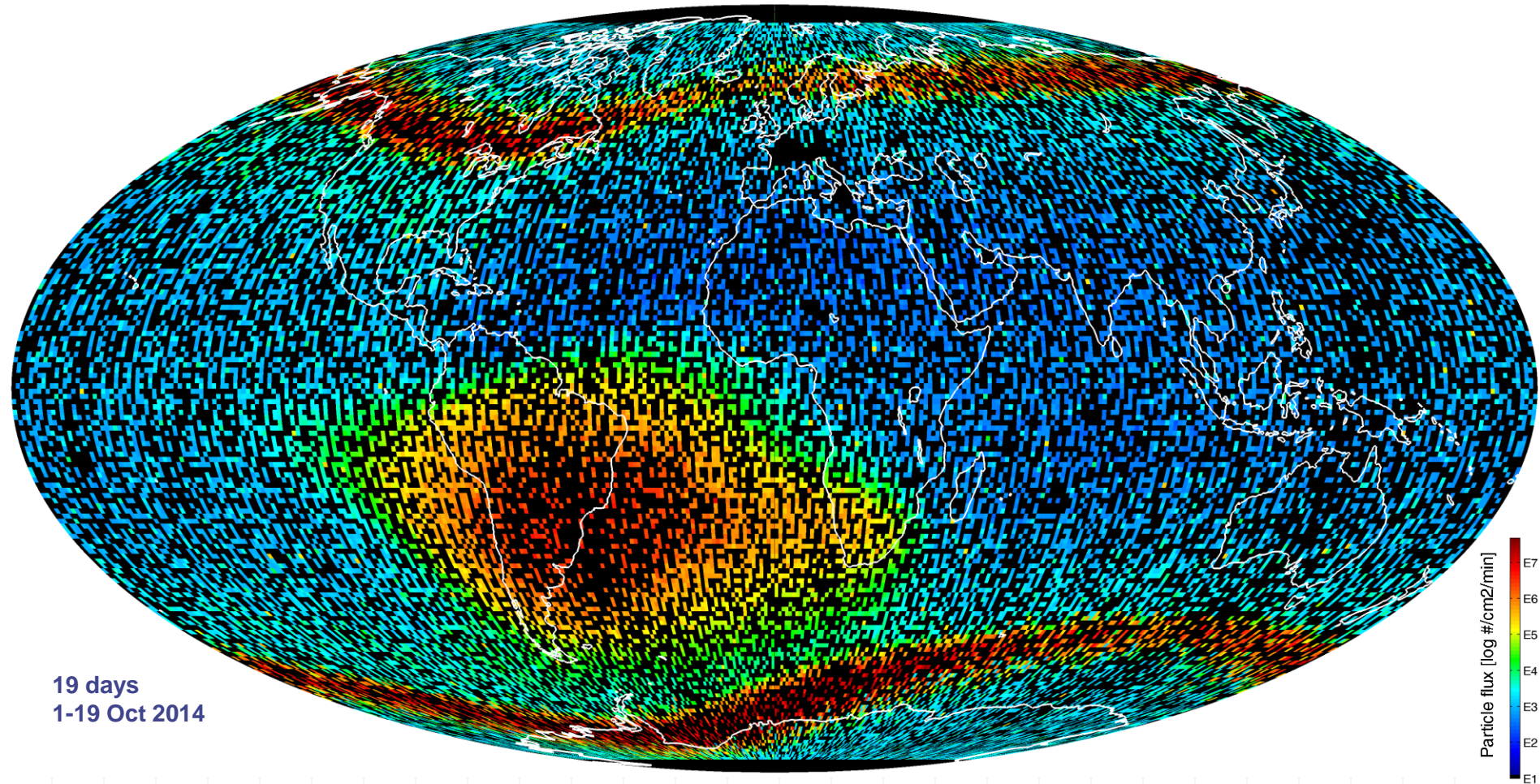
- Quantum imaging detection
- Resolving power radiation components
- directional sensitivity
- dE + track path → LET

Timepix SATRAM/ESA Proba-V in Open Space

Quantum imaging detection/monitoring of space radiation



Map: particle flux (all particles)

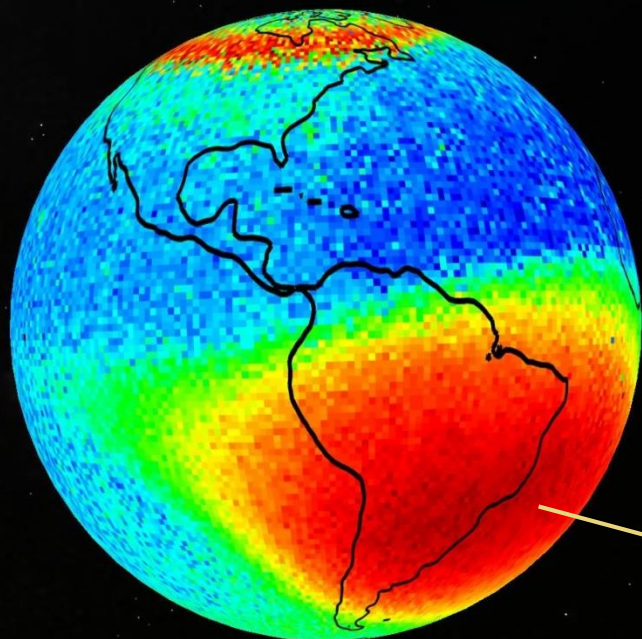
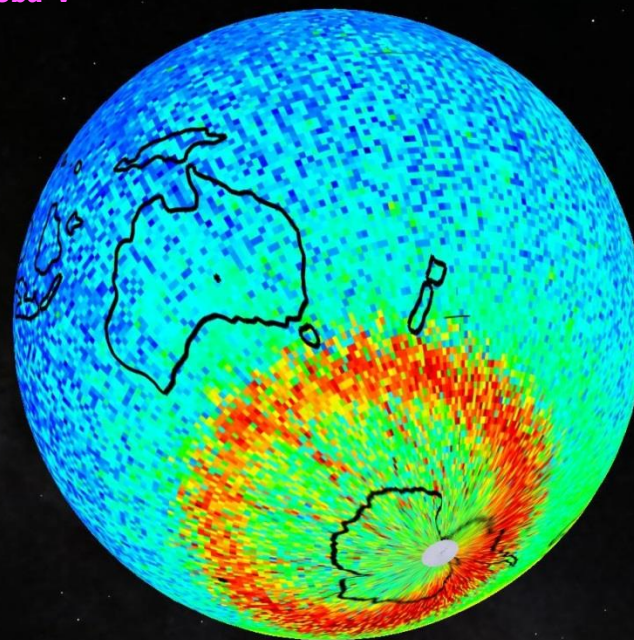
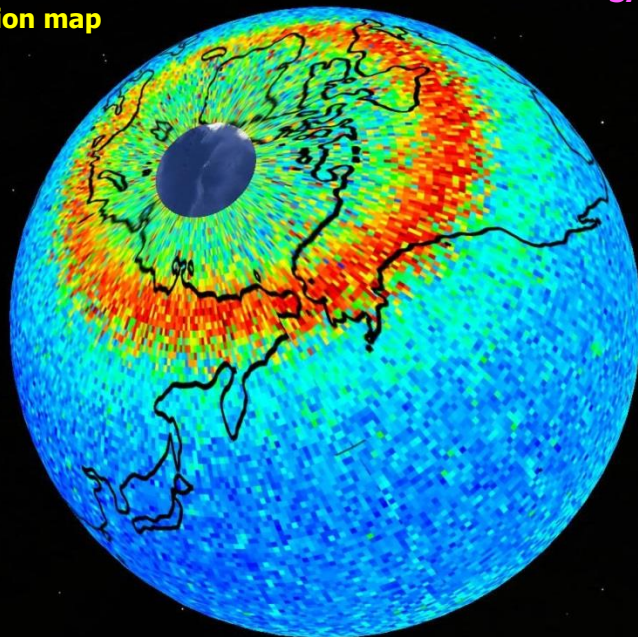


19 days
1-19 Oct 2014

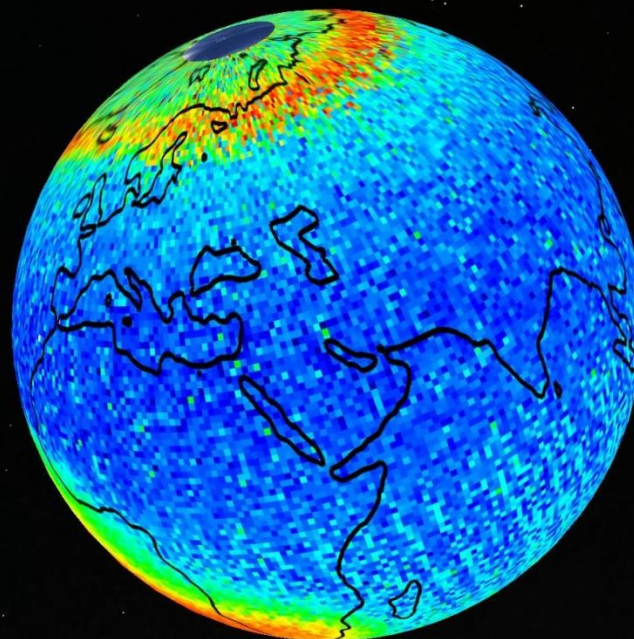
Particle flux [log #/cm²/min]
E7
E6
E5
E4
E3
E2
E1

Space radiation map
at LEO orbit
all particles

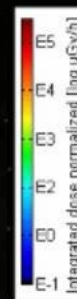
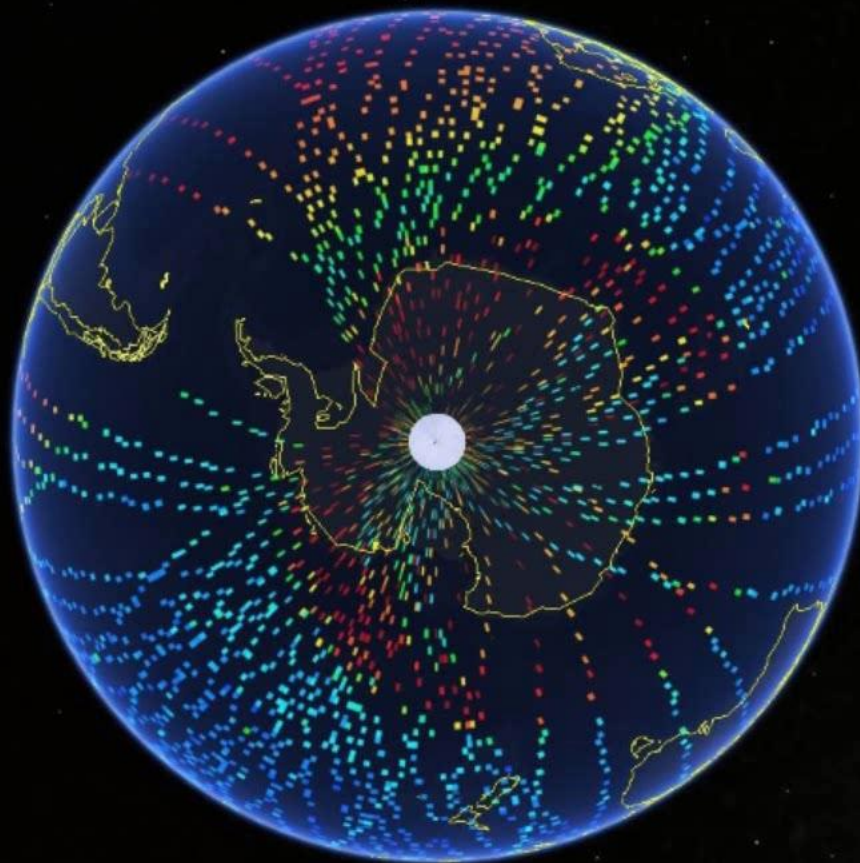
SATRAM Timepix/Proba-V



SAA

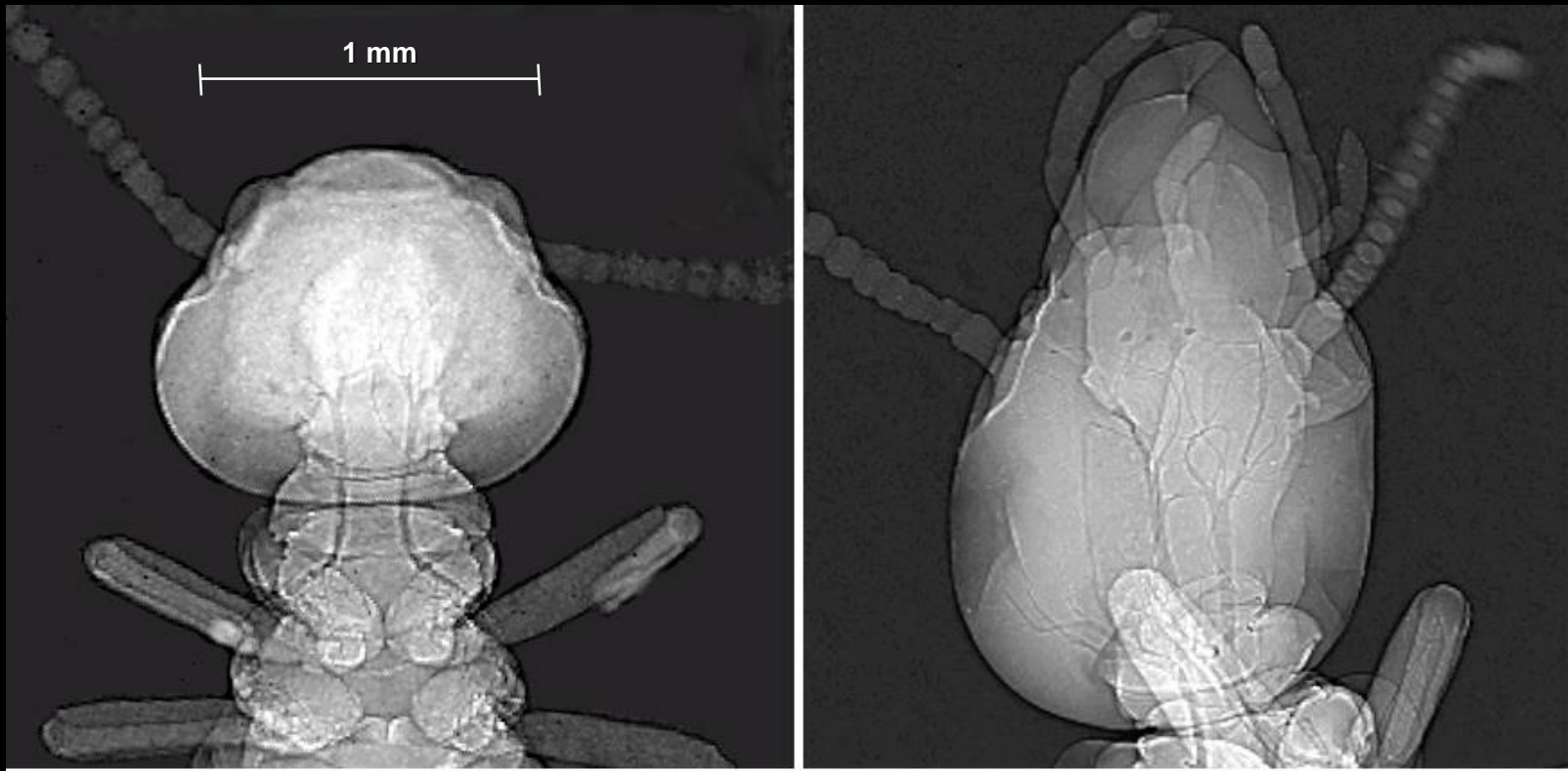


3-9 Jan 2014



High resolution high contrast X-ray imaging

Living termites + observation of dynamic biological processes



*Thanks
to the
organizers +
audience*

- X-ray radiographs of a termite worker before (left) and after 20 h (right) its metamorphosis toward the soldier caste (5s exposure ~ 0.7mGy dose)
- Micro-focus X-ray tube + Timepix detector
- Phase-contrast enhanced sensitivity

