





#### X-Ray Fluorescence Imaging with a Laser-Wakefield Thomson X-Ray Source

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PLASMED X

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- Principle of X-ray fluorescence imaging (XFI)
- Requirements on the X-ray source & necessary improvements
- Laser-wakefield acceleration (LWFA)
- Thomson scattering (TS)
- First proof-of-principle experiment and results
- Optimisation of the setup



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# X-RAY FLUORESCENCE IMAGING (XFI)

- Incident X-ray beam displaces electrons from inner orbital shells of the target atom
- Vacancies are filled by electrons from higher orbits
- Energy difference is released in form of characteristic X-rays
- Potential use for medical imaging with high-Z elements



Taken and modified from www.bruker.com



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# PREVIOUS IMAGING OF GNPs IN A TUMOR- BEARING MOUSE

- Mouse injected with small amounts of gold nanoparticles (GNPs)
- Benchtop setting with polychromatic X-ray source
- Detect characteristic gold fluorescence signals
- Translation of the detector in order to obtain tomographic images





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Scientific Reports 2018, 8:16561, F.Grüner et. al.

# LOCALISING FUNCTIONALISED GOLD-NANOPARTICLES

- GNPs functionalised with L1-peptides injected into murine spinal cord → bind to stressed neurons
- XFI-scans localise regions with bound GNPs
- Gold masses down to 72 pg could be detected
- Close agreement to inductively coupled plasma mass spectrometry (ICP-MS) results



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#### NECESSARY IMPROVEMENTS & REQUIREMENTS

- Surface-functionalisation of GNPs
- Development of large-area, pixelated, spectroscopic detectors
- Development of compact, hard X-ray sources (80 100 keV) with high brilliance, flux and repetition rate
- Small source size
- Inexpensive realisation
- Effective background reduction with advanced filtering scheme and collimators



Pixelated Hexitec detector from www.quantumdetectors.com

Antibodies Peptides SiRNA Aptamers

> Diagram of surface functionalisation of nanoparticles from www.moleculardevices.com



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#### CLINICAL SOURCES TODAY: X-RAY TUBES

bremsstra hlung

- Electrons are discharged and accelerated
- Collisional and radiative energy transfer
- Bremsstrahlung and characteristic X-rays
- 1% of the energy : X-rays
- 99% of the energy: heat





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#### SYNCHROTRONS

- Modern synchrotrons provide high energy and brilliance photons
- Circumferences in the order of kilometers and large numbers of bending magnets
- $\rightarrow$  huge and expensive
- Not suitable for medical applications (except at research centers)

Aerial view of the PETRA III synchrotron at DESY in Hamburg (http://photon-science.desy.de)



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#### LASER-WAKEFIELD ACCELERATION (LWFA)





#### FAKULTÄT

FÜR MATHEMATIK, INFORMATIK UND NATURWISSENSCHAFTEN DESY Strategy Fund.



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#### LAB SETUP @DESY/HAMBURG





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# THOMSON SCATTERING (TS)



Simple schematic of the Thomson scattering process





Photon number:  $N_{\gamma} \sim Q_{bunch} \tau_{laser} a_0^2 \sigma(\theta_{obs})$ 

EAAC WG4, 18.09.19, theresa.staufer@desy.de



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**THOMSON SCATTERING X-RAY SOURCES** incoming laser beam parabola focusing mirror beam splitter main laser pulse Thomson scattering laser focusing parabola Inverse Compton scattering electrons transverse probe beam gas jet X-rays beamsplitter gas jet Object / Detector Schematic of the Thomson scattering setup Sketch of the Thomson spherical mirror scattering setup in the vacuum chamber electrons and X-rays exit the differential chamber pumping system



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#### FIRST PROOF-OF-PRINCIPLE EXPERIMENT



- Hexitec detector<sup>1</sup> at 8 meters distance from the interaction point
- 5 minutes measurement duration
- 46% laser power (2-3 pC charge, 60 MeV mean electron energy)
- Clear difference with / without
  Thomson scattering laser
  - $^{\rm 1}$  kindly loaned from CLF



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#### IMAGING OF GNPs IN A MOUSE PHANTOM



- 3 cm diameter PMMA-cylinder with 0.3 ml Eppendorf tube containing **10 mg/ml** GNPs
- Target at 3.25 meters from interaction point
- Calculated significance **Z** = **5.4**





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#### IMAGING OF GNPs IN A MOUSE PHANTOM



- 3 cm diameter PMMAcylinder with 0.3 ml
   Eppendorf tube containing 18.87 mg/ml GNPs
- Improved lead shielding to reduce
  - Bremsstrahlung
- Target at 3.25 meters from interaction point
- Calculated significanceZ = 8.3



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# IMAGING OF Gd-SOLUTION

- Gadolinium is used as contrast agent in magnetic resonance imaging
- 1 cm diameter Eppendorf tube containing **78 mg/ml** Gd-solution
- Target (only Eppendorf tube, no surrounding phantom) at 3.25 meters from interaction point
- Calculated significance of fluorescence lines at 42.3 and 43 keV: Z = 32
- Even  $K_{\beta}$ -fluorescence at 48.7 keV visible





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# **OPTIMISATION OF THE SETUP**

- Improving the electron parameters:
  - **Active Plasma Lenses**
- Improving the laser parameters: maximising X-ray gain by varying laser focus and length
- Simulated result: 80 000 photons per shot in **90 keV ± 15 % FWHM** (ideal parameters for XFI)



Schematic of the optimised setup with an active plasma lens



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#### SUMMARY

- XFI is a new medical imaging modality with applications e.g. in pharmacokinetics or tumor diagnostics
- Developments in accelerator technologies enable to build compact and high-brightness X-ray sources
- Thomson scattering with laser-wakefield acceleration is an excellent driver for XFI with desired properties
- Proof-of-principle experiments demonstrated XFI measurements with a laser-driven Thomson X-ray source
- Implementation of active plasma lens will optimise incident Thomson spectrum for XFI by reducing effective spectral width of electrons