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# Advanced X-ray imaging: Spectral and Phase-contrast Techniques

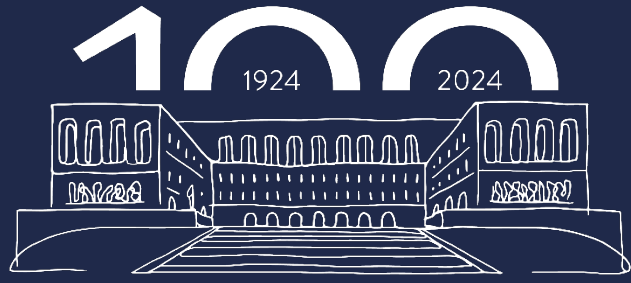
Workshop on Particle Detectors for Interdisciplinary Applications  
Dept. of Physics, University de los Andes, Bogota, Colombia (online)  
October 1<sup>st</sup>, 2024

Luca Brombal

Universidad de los Andes | Facultad de Ciencias  
Departamento de Física

2024

Workshop on Particle Detectors for  
Interdisciplinary Applications.  
September 30 – October 2.



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# Financial interests and relationship

**NOTHING TO DISCLOSE!**

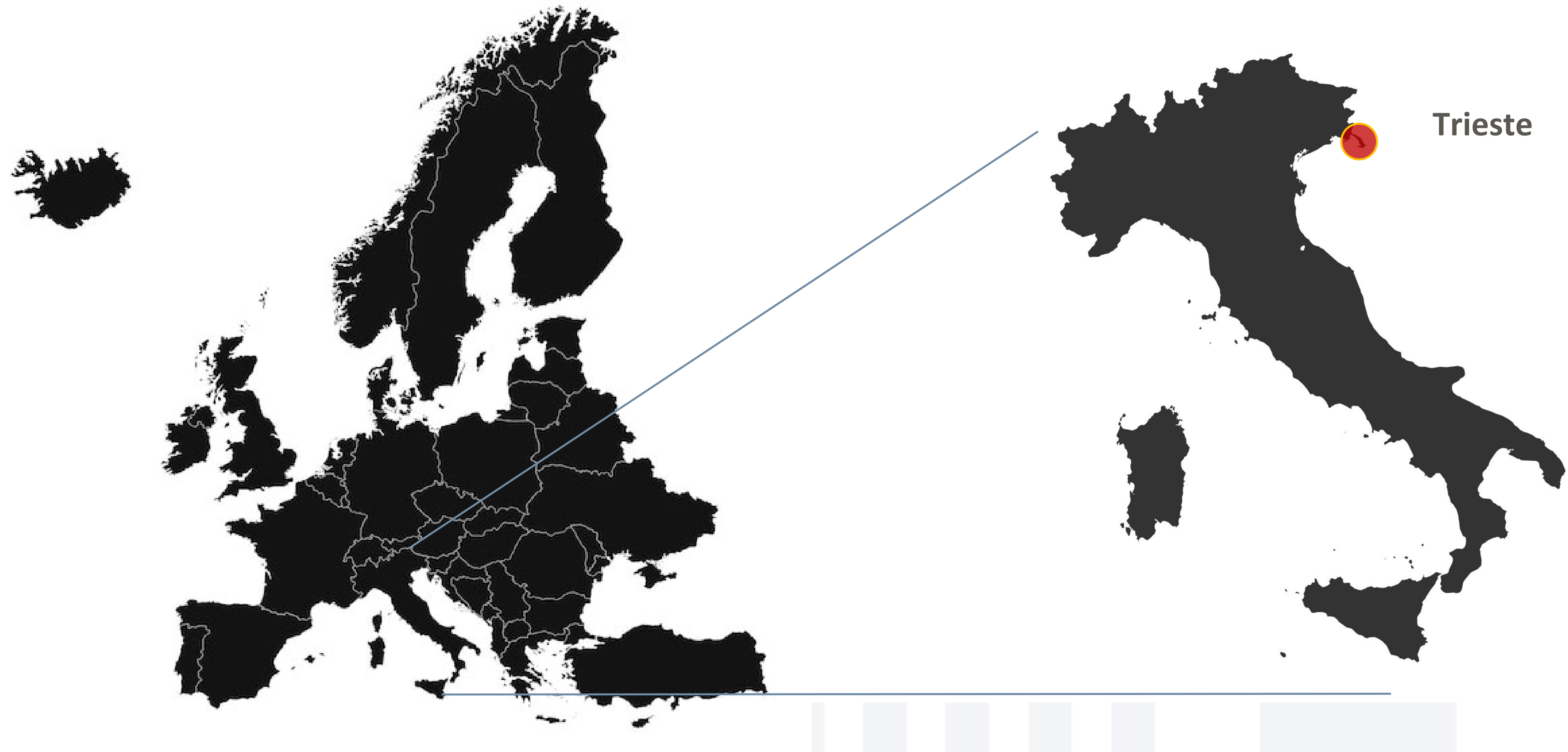
## Funding

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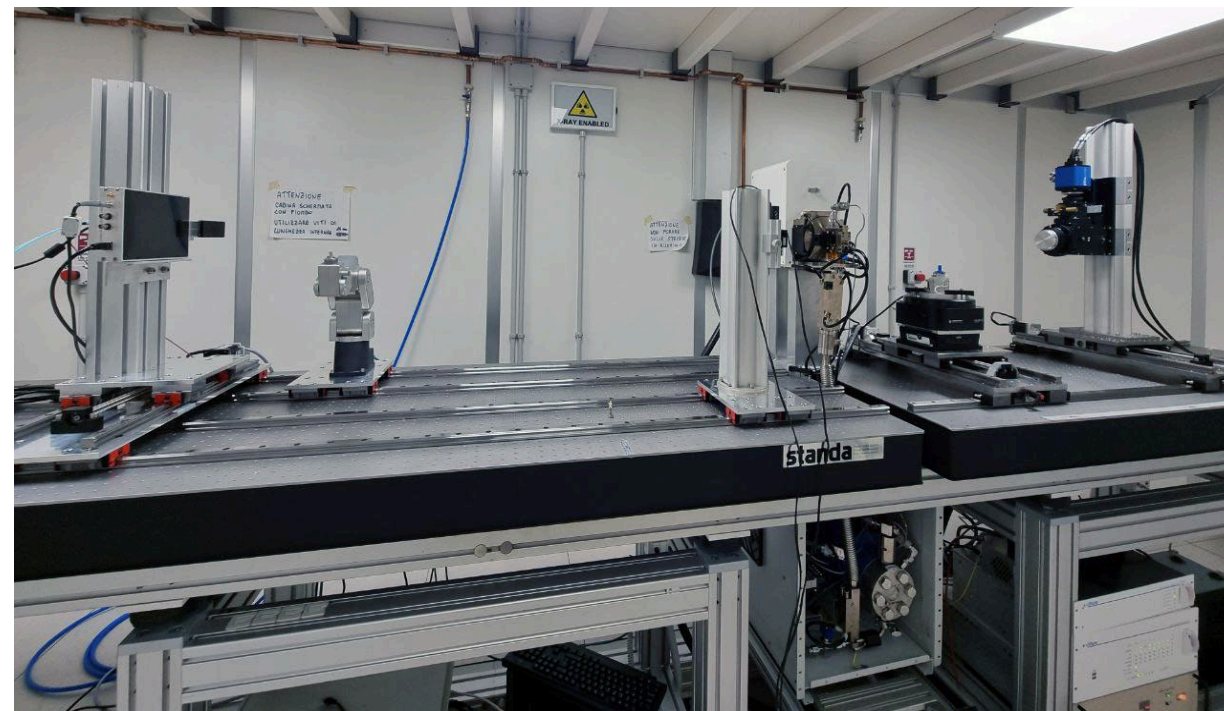
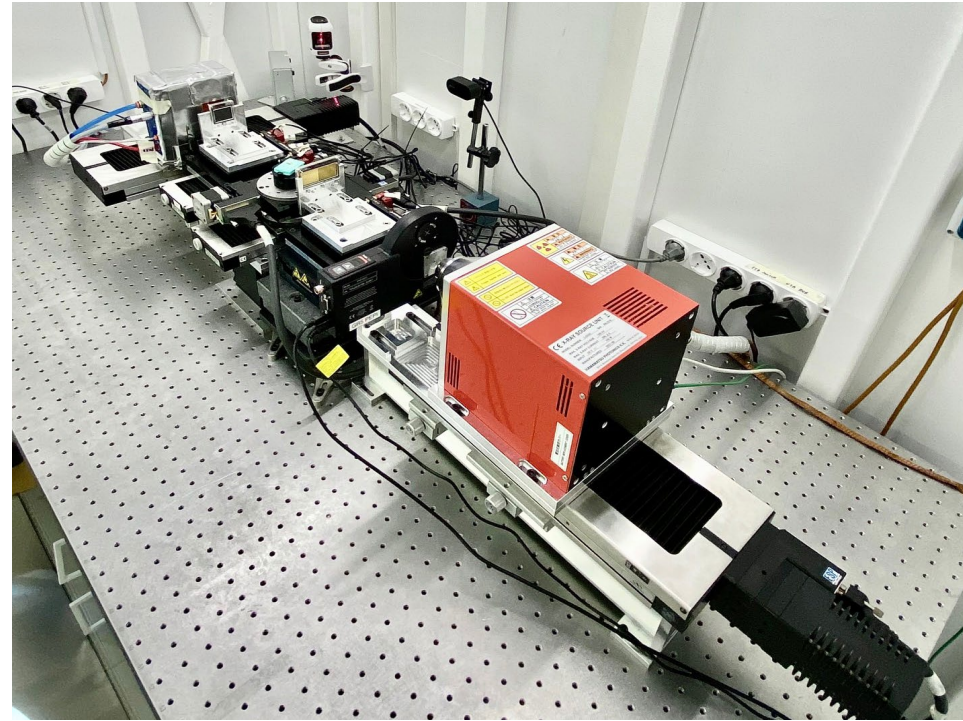
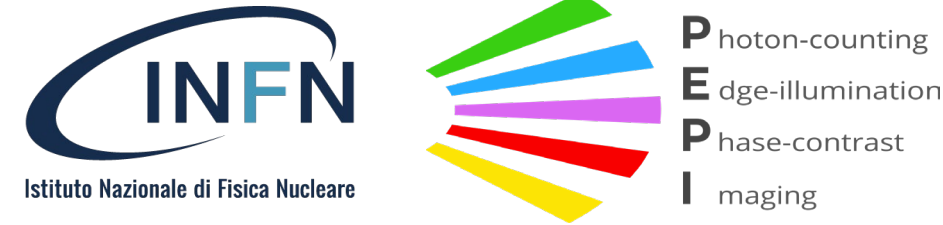


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# TRIESTE X-RAY TOMOGRAPHY COLLABORATIVE



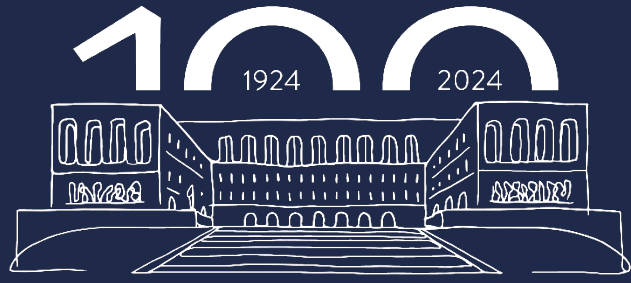
# TRIESTE X-RAY TOMOGRAPHY COLLABORATIVE



OPTIMATO LAB



SYRMEP BEAMLINE + TOMOLAB



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# OUTLINE

- **X-ray imaging fundamentals**
- Spectral imaging
- Phase-contrast imaging
- Spectral phase-contrast imaging

# WHAT'S CONVENTIONAL X-RAY IMAGING?

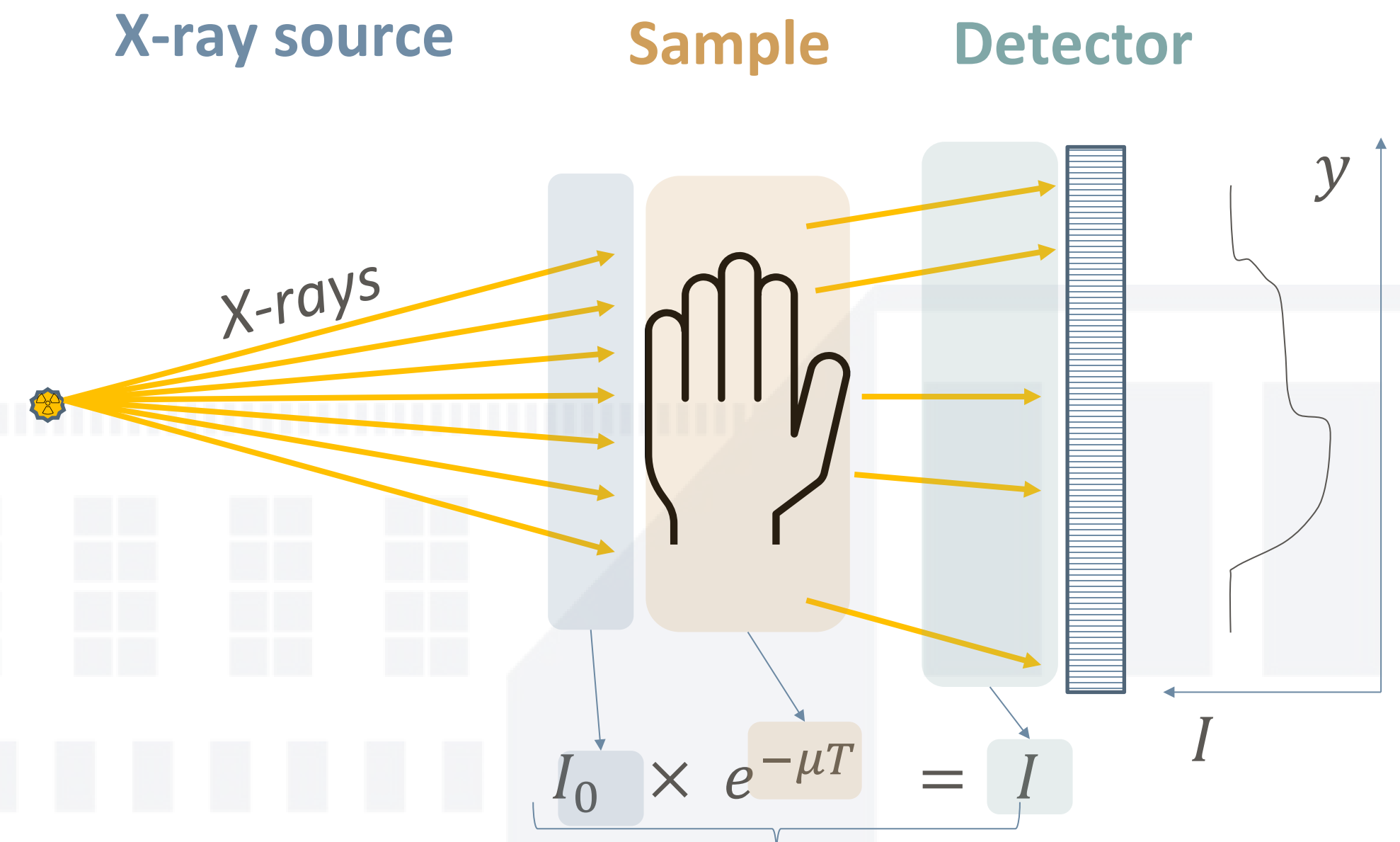
The basics before getting advanced...

The key elements of conventional X-ray imaging are:

1. **X-ray source** (e.g., X-ray tube)
2. **Sample** to be investigated
3. **Detector** sensitive only to X-ray intensity

Sample's visibility depends on the (partial) attenuation of X-rays:

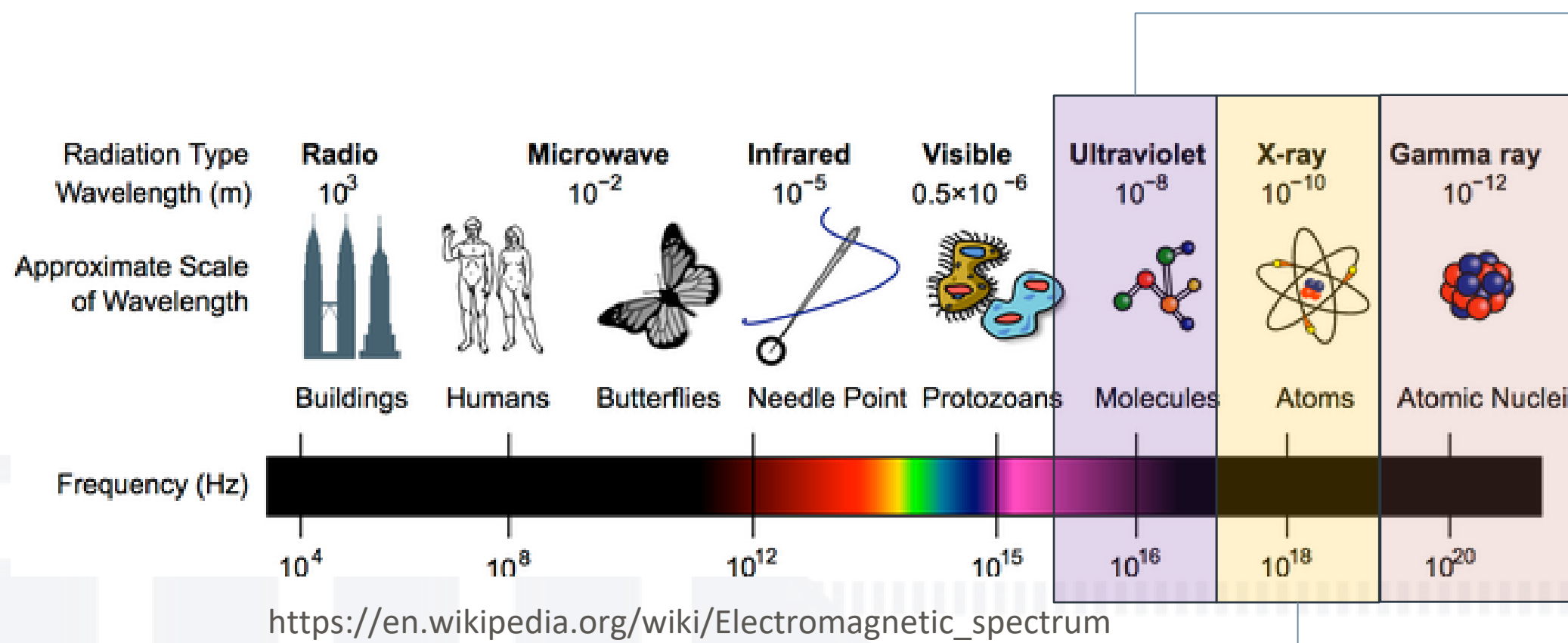
- Sample thickness ( $T$ )
- Linear attenuation coefficient of the sample ( $\mu$ )



*Beer-Lambert's law*

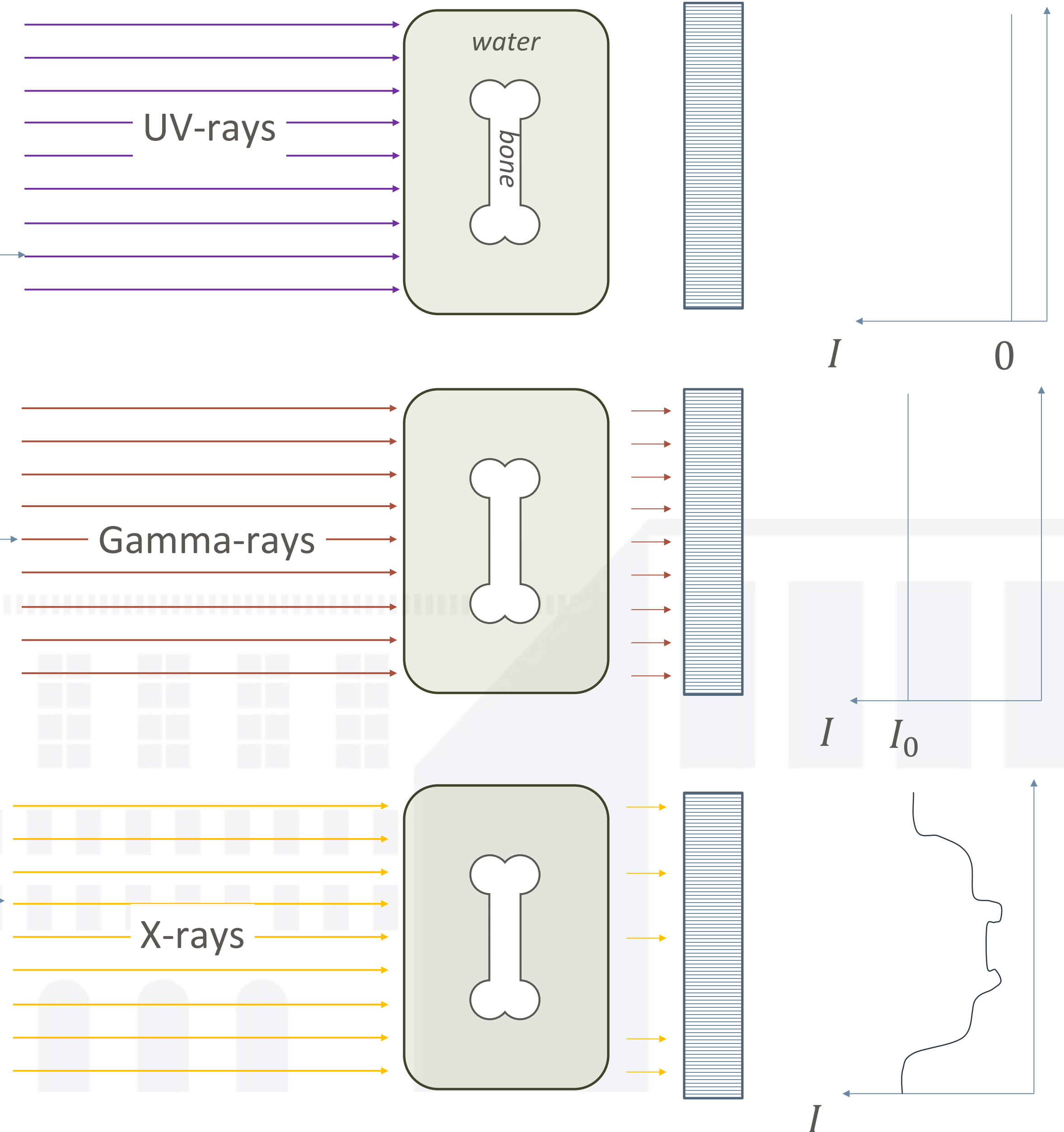
# WHY X-RAYS?

The basics before getting advanced...



Because:

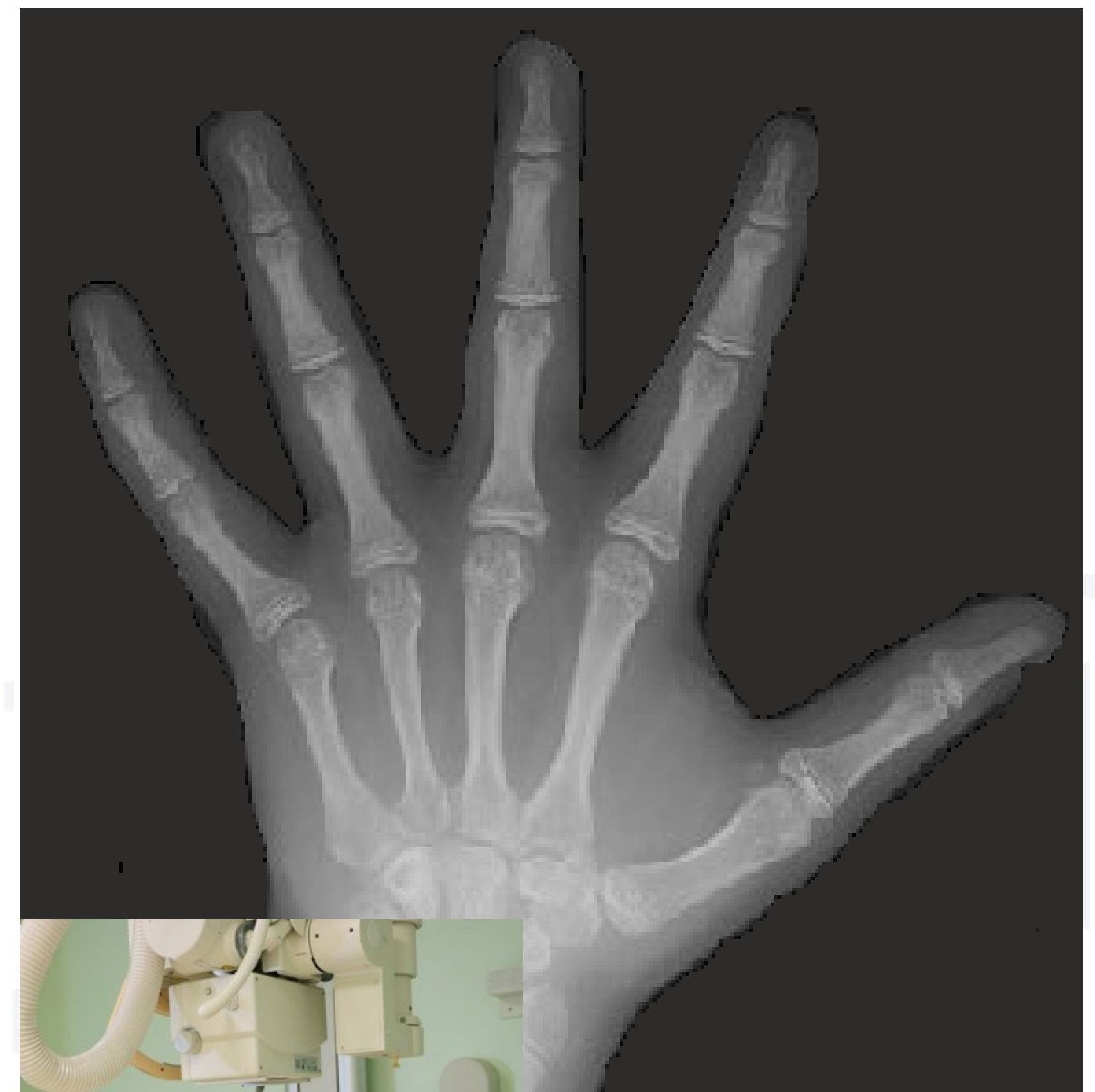
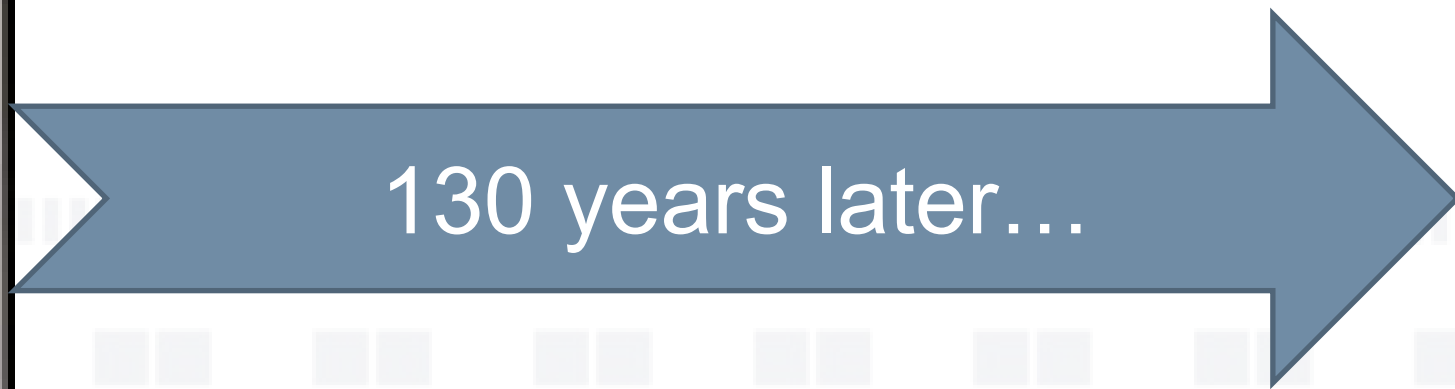
1. They have a **sufficiently weak interaction** with matter → they can partly go through
2. They have a **sufficiently strong interaction** with matter → contrast (i.e. attenuation difference) among transversed features is formed on the detector



# 130 YEARS OF PROGRESS...



**First radiograph by  
Wilhelm Conrad Roentgen**  
- Fall 1895,  
- 1<sup>st</sup> Nobel Prize in Physics 1901



← **First commercial device  
spring 1896**

Commercial devices →  
nowadays

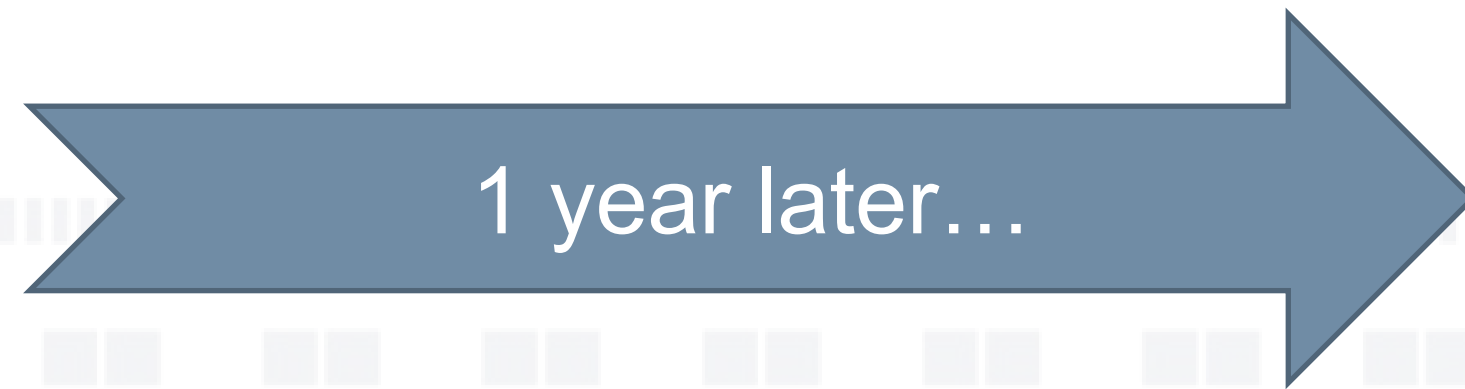




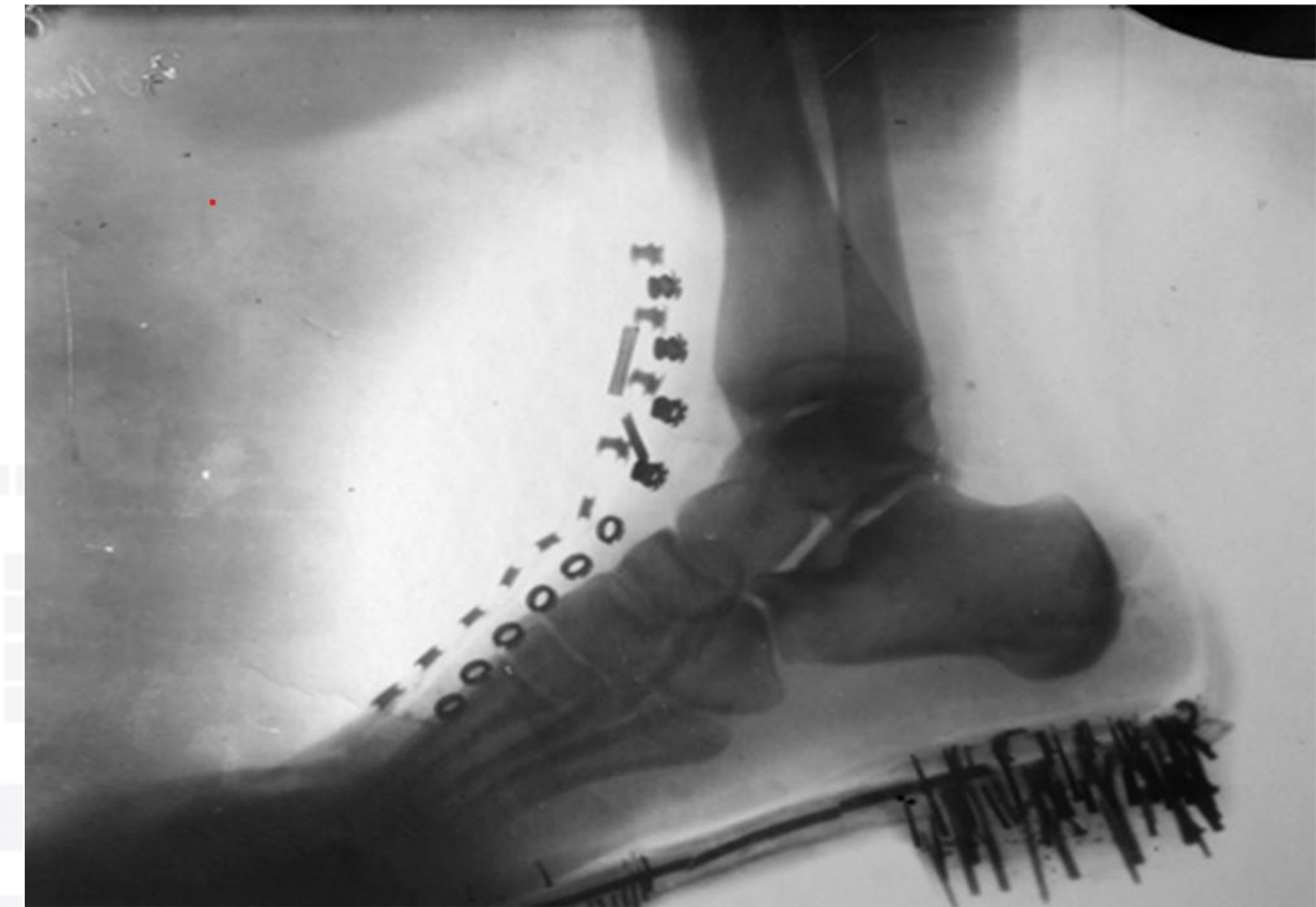
# ...CONSPIRACY THEORY DETOUR...



**Wilhelm Conrad Roentgen**  
Fall 1895



**Nikola Tesla**  
1896



**Was Roentgen the first to discover X-rays?**  
**Check this out**

<https://doiserbia.nb.rs/img/doi/0025-8105/2016/0025-81051610313V.pdf>

# ADVANCED X-RAY IMAGING

## CONVENTIONAL IMAGING

1. X-ray source – conventional X-ray tube
2. Detector sensitive only to beam intensity
3. Sample to be investigated



## ADVANCED IMAGING

1. X-ray source with high coherence (spatial/temporal) or **capable of producing different X-ray spectra at the same time**
2. **Detector sensitive to the energy spectrum of x-rays**
3. Optical elements to condition the beam upstream and/or downstream of the sample

**SPECTRAL  
imaging**

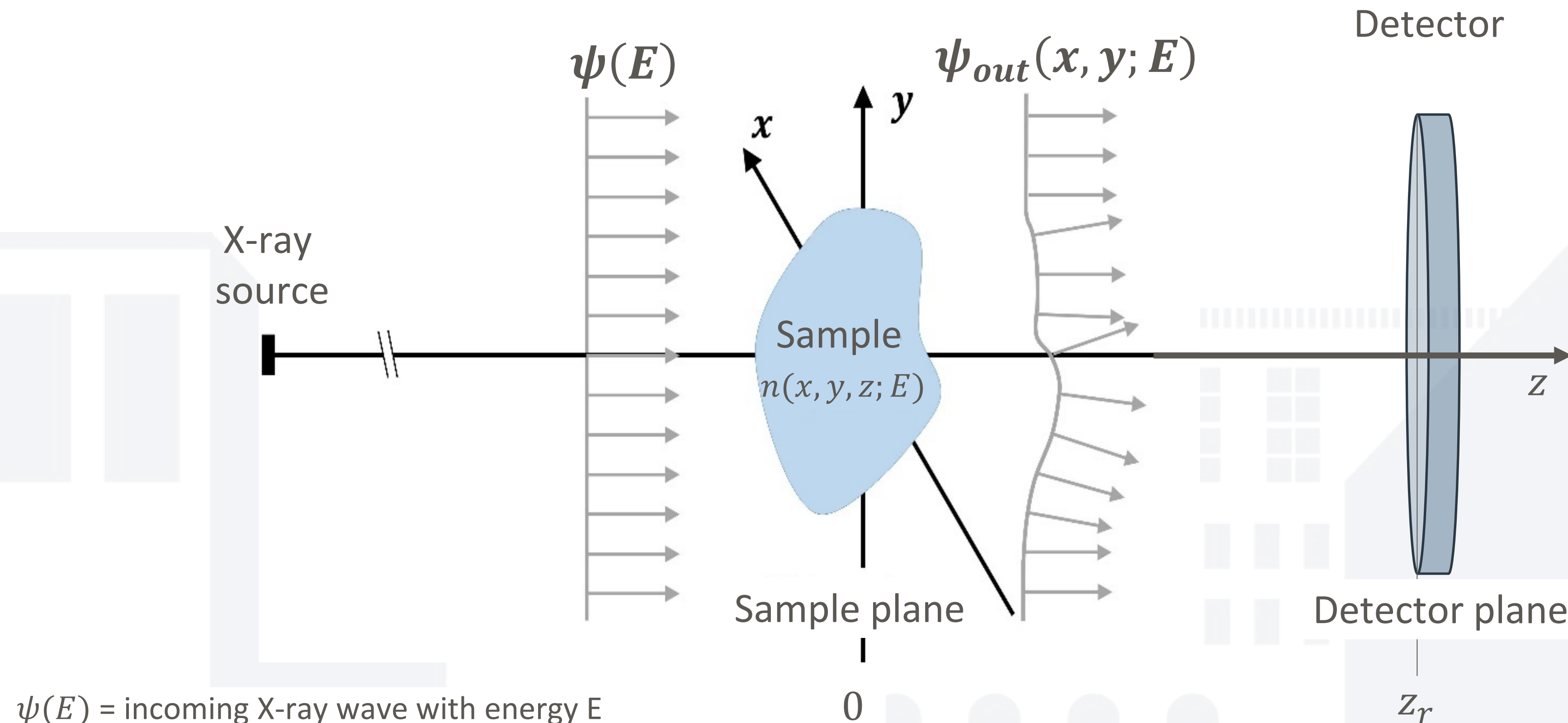
**PHASE-  
CONTRAST  
imaging**

## ADVANCED IMAGING

1. X-ray source with high coherence (spatial/temporal) or **capable of producing different X-ray spectra at the same time**
2. **Detector sensitive to the energy spectrum of x-rays**
3. Optical elements to condition the beam upstream and/or downstream of the sample

# X-RAY IMAGING IN WAVE FORMALISM

Just a few slides



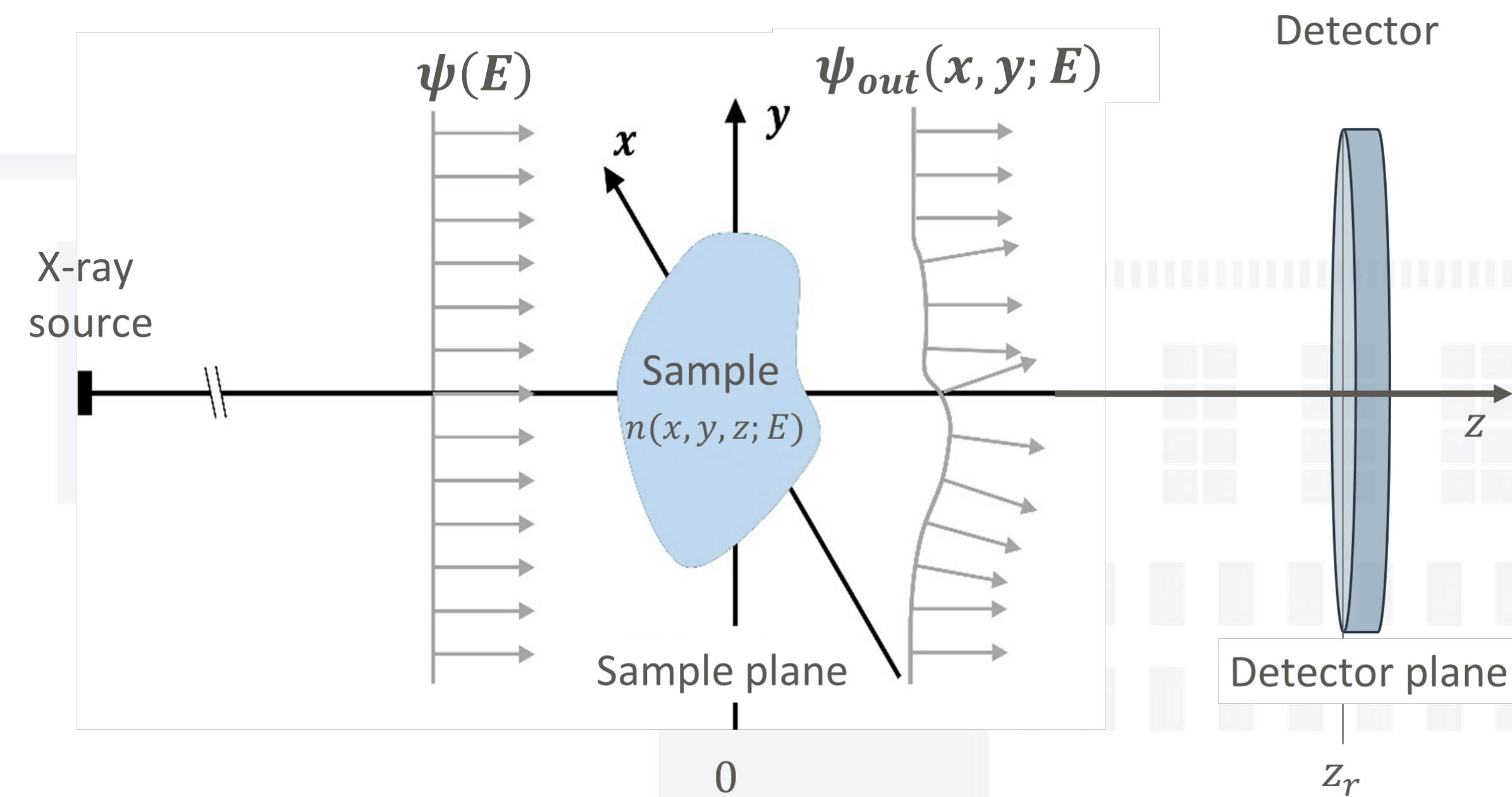
$\psi(E)$  = incoming X-ray wave with energy  $E$   
 $\psi_{out}(x, y; E)$  = outgoing X-ray wave modulated by the sample  
 $n(x, y, z; E)$  = complex refractive index

*Rigon, Luigi. "X-ray imaging with coherent sources." (2014): 193-216.*

# X-RAY IMAGING IN WAVE FORMALISM

Just a few slides

- In the wave model the interaction of X-rays with matter is described through the complex refractive index  $n$



$$n(E) = 1 - \delta(E) + i\beta(E)$$

$$\beta(E) = \frac{\mu(E)}{2k}$$

absorption coefficient

$$\psi_{out}(x, y; E) = \psi_{in}(E) e^{-k \int \beta(x, y, z; E) dz} e^{-ik \int \delta(x, y, z; E) dz}$$

Attenuation

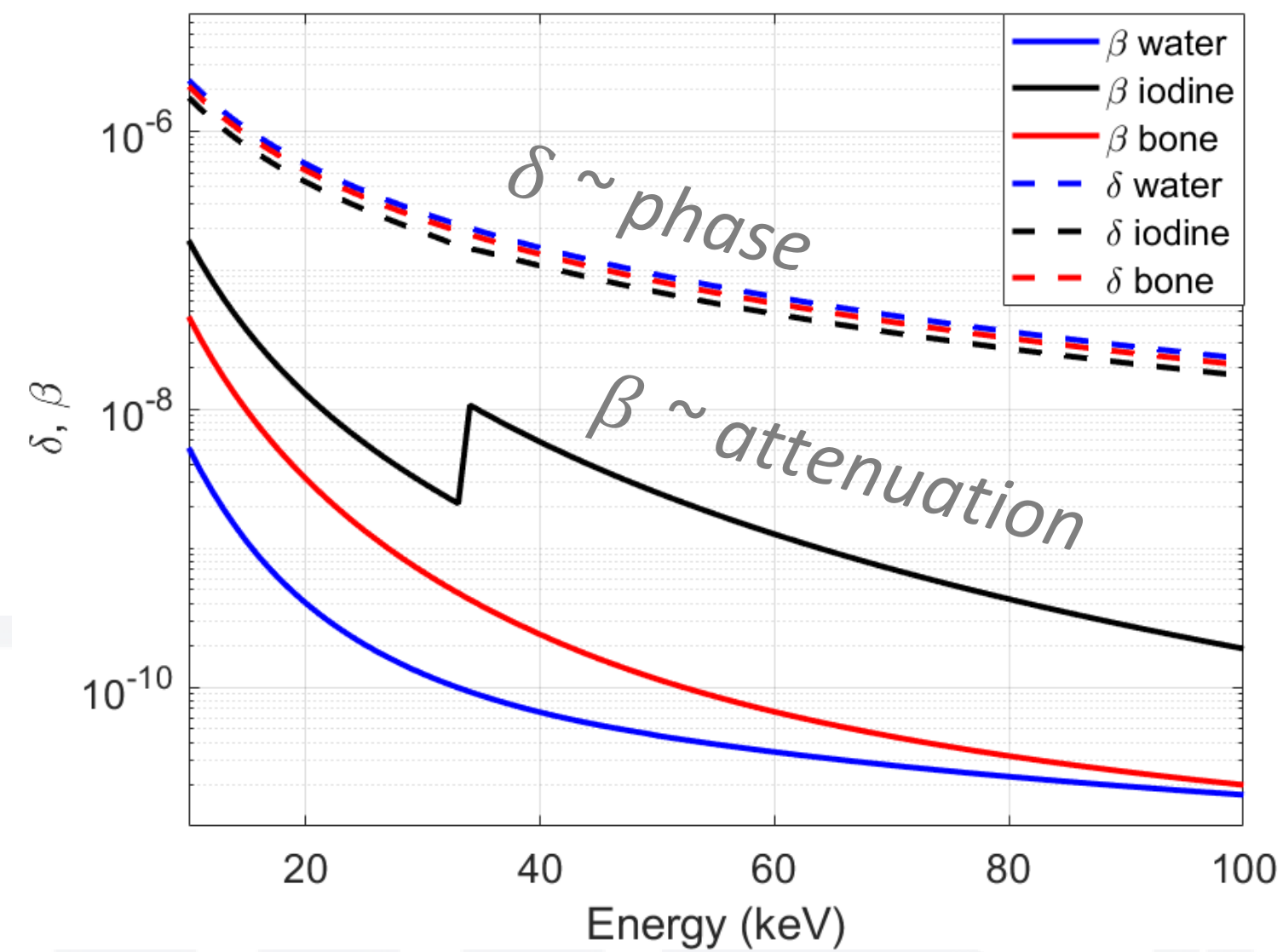
Phase shift

Spectral imaging

Phase-contrast imaging

$\psi_{in}(E)$  = incoming plane wave at energy  $E$   
 $\psi_{out}(x, y; E)$  = object-modulated wave at energy  $E$   
 $k$  = wave number

# WHY SPECTRAL?



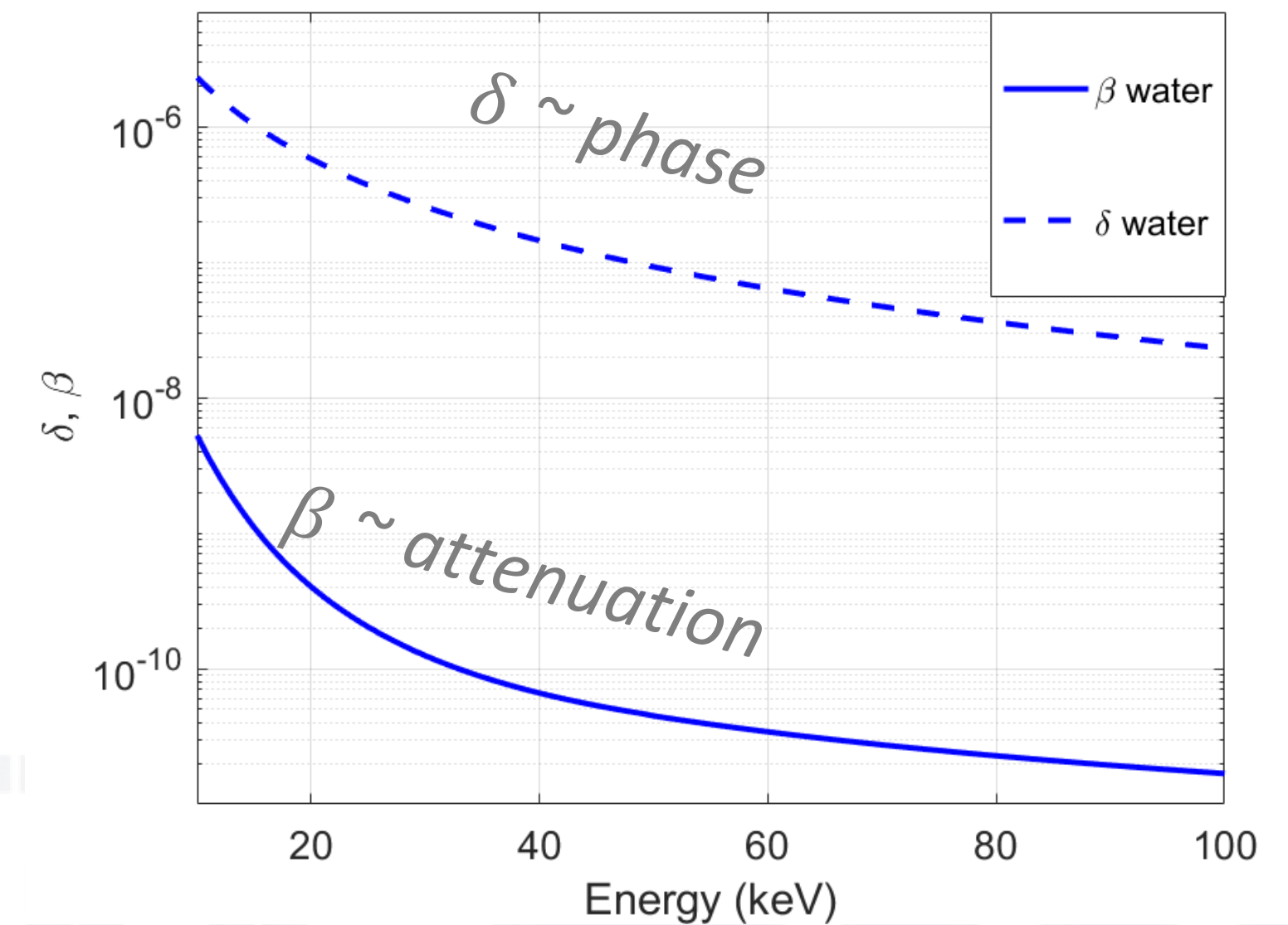
- Energy dependence of the attenuation coefficient is sensitive to chemical composition (i.e., difference in atomic number  $Z$ )

$$\mu(E) = \mu_{\text{photoelectric}}(E) + \mu_{\text{scattering}}(E) + \mu_{K_{\text{edge}}}(E)$$

$$\propto \frac{Z^4}{E^{3.5}} \quad \propto Z \quad \neq 0 \text{ at } K_{\text{edge}}$$

to separate and quantify chemical elements!

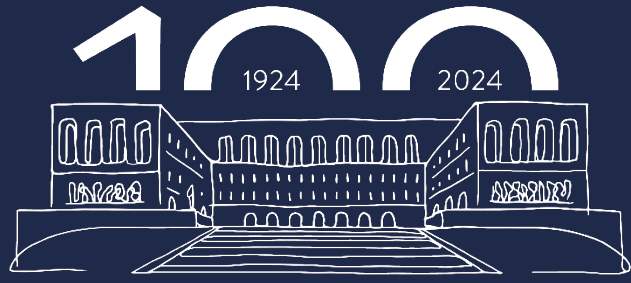
# WHY PHASE-CONTRAST?



- For light materials and energies in the range 10 – 100 keV phase effects are much larger than attenuation

$$\frac{\delta}{\beta} \sim 10^3$$

to increase visibility of light materials!  
(e.g. soft tissues)



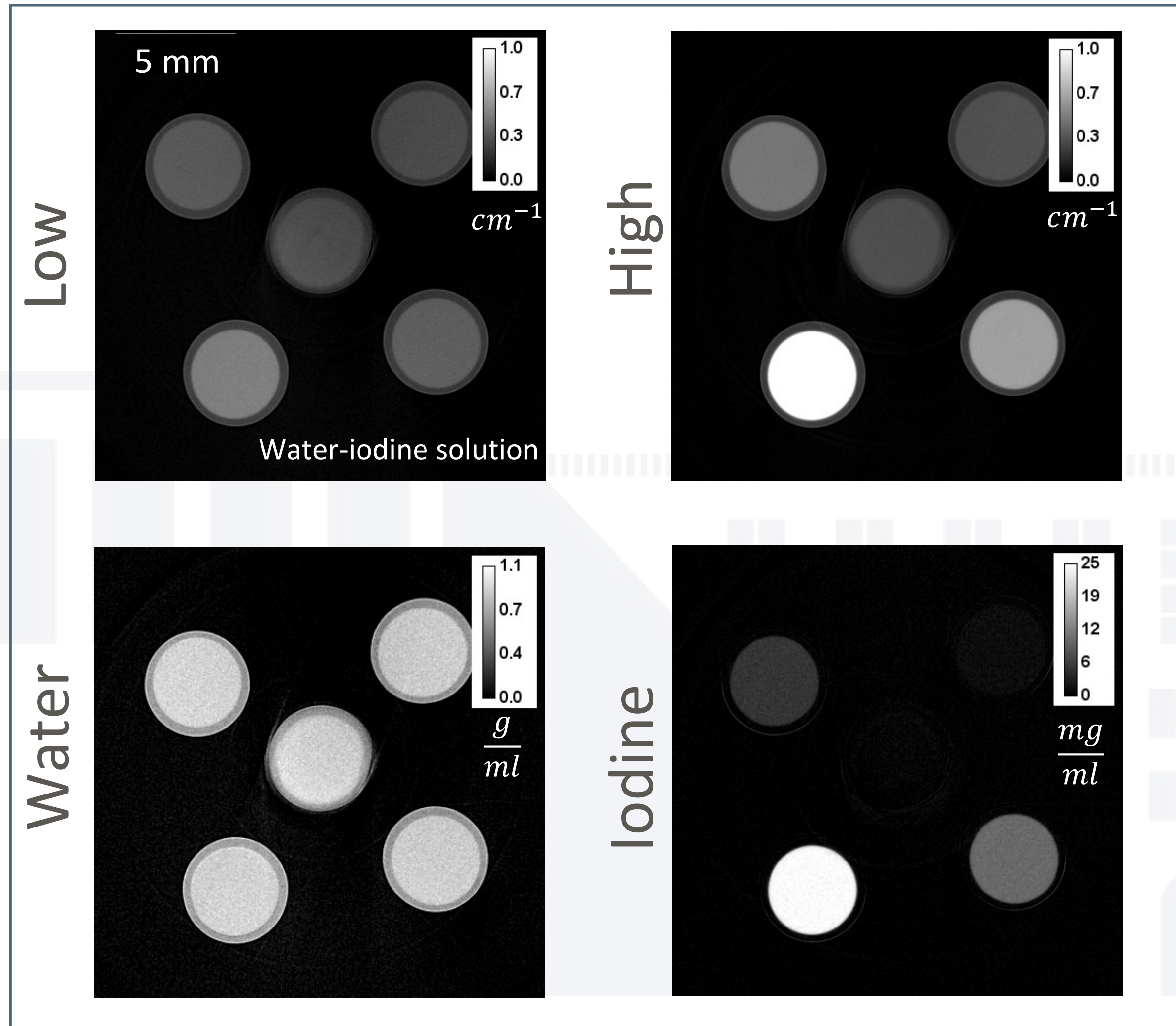
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# OUTLINE

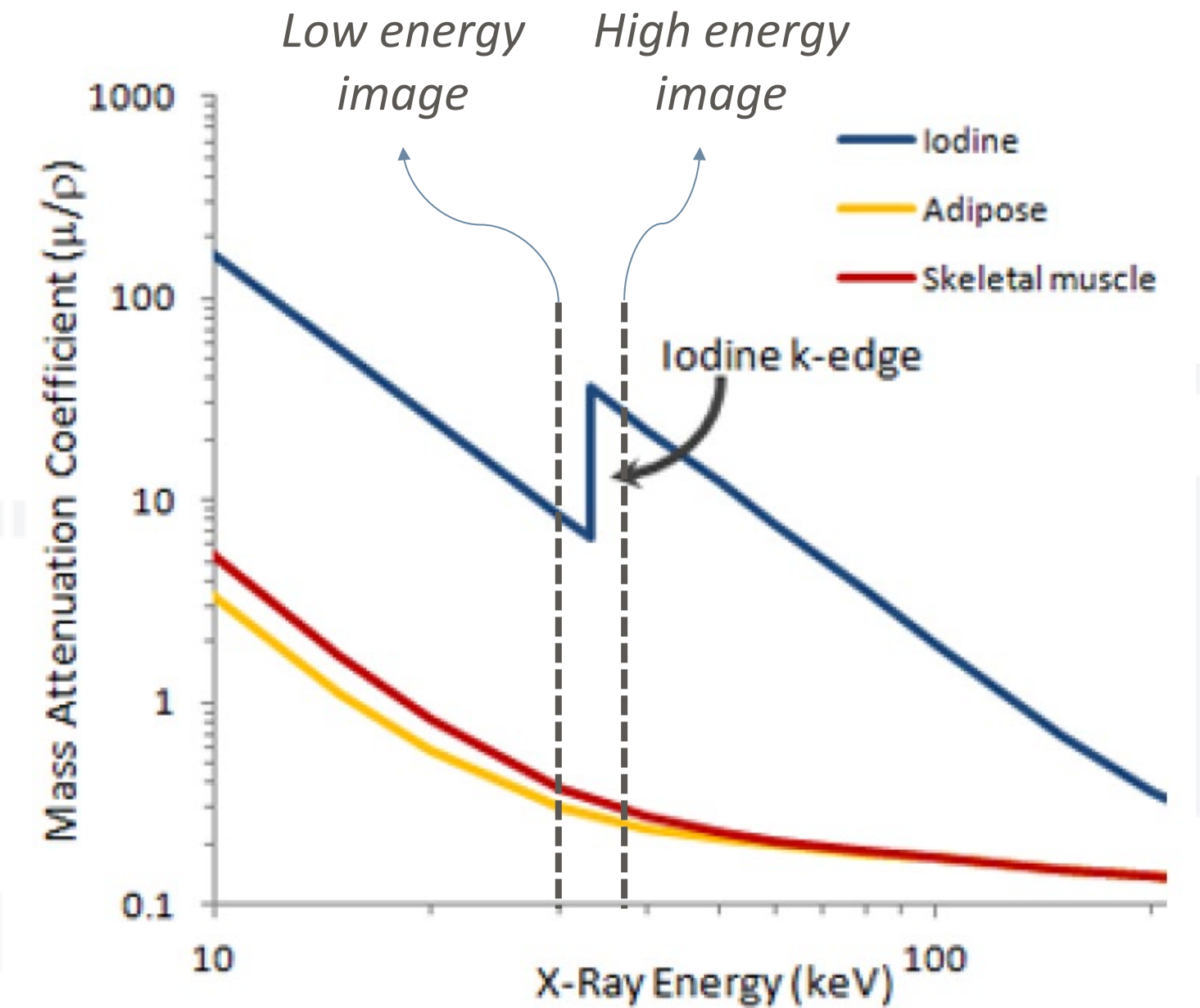
- X-ray imaging fundamentals
- **Spectral imaging**
- Phase-contrast imaging
- Spectral phase-contrast imaging

# X-RAY SPECTRAL IMAGING

Acquired @ Elettra – bent Laue crystal



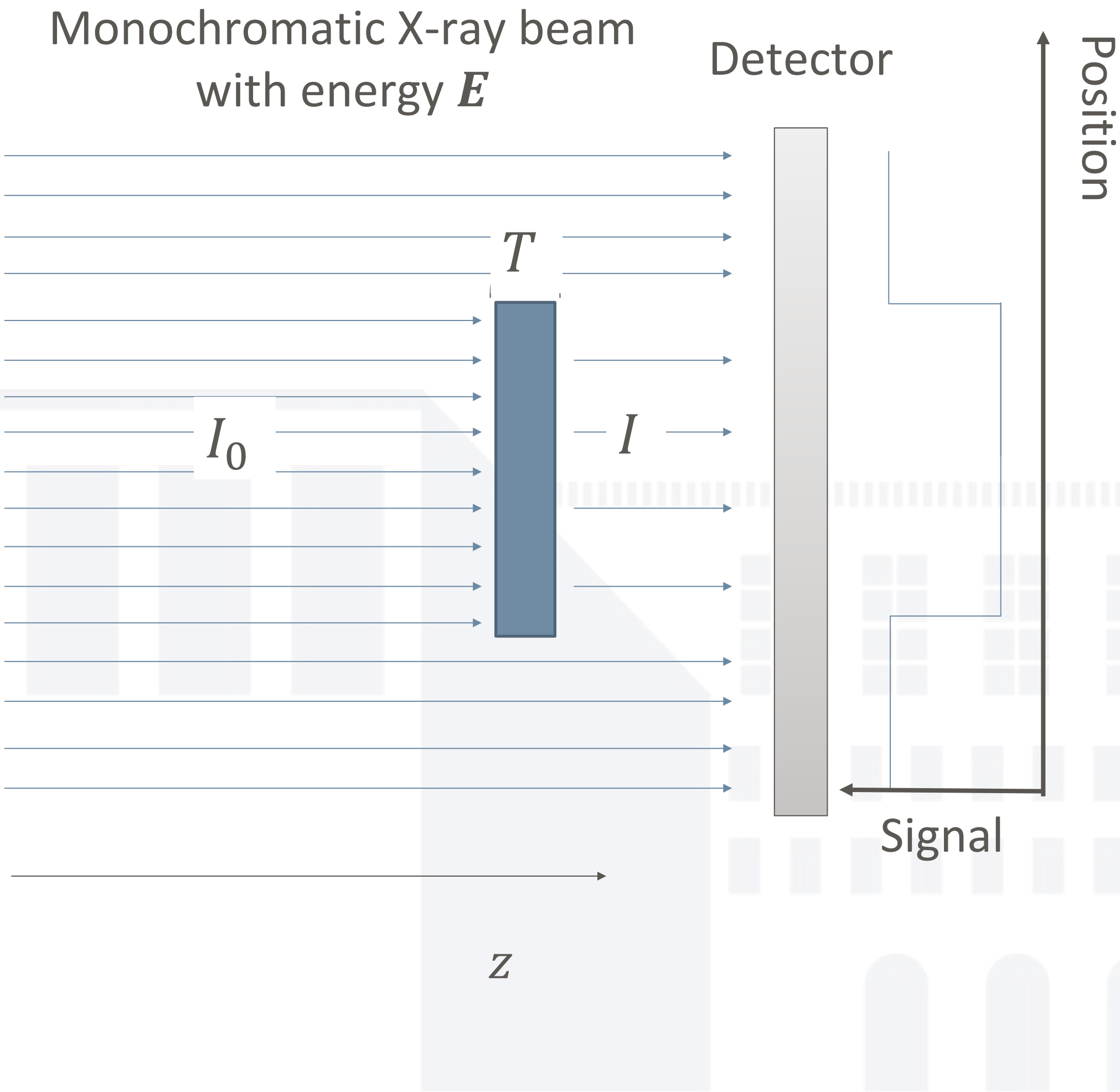
- Spectral imaging requires to probe the attenuation properties of the sample (at least) at 2 different energies



- Images acquired at different energies are processed through matrix inversion algorithms to **extract (quantitative) maps of elements of interest**



# X-RAY SPECTRAL RADIOGRAPHY – BASICS (1)



Beer-Lambert law

$$I = I_0 e^{-\int \mu(E) dz}$$

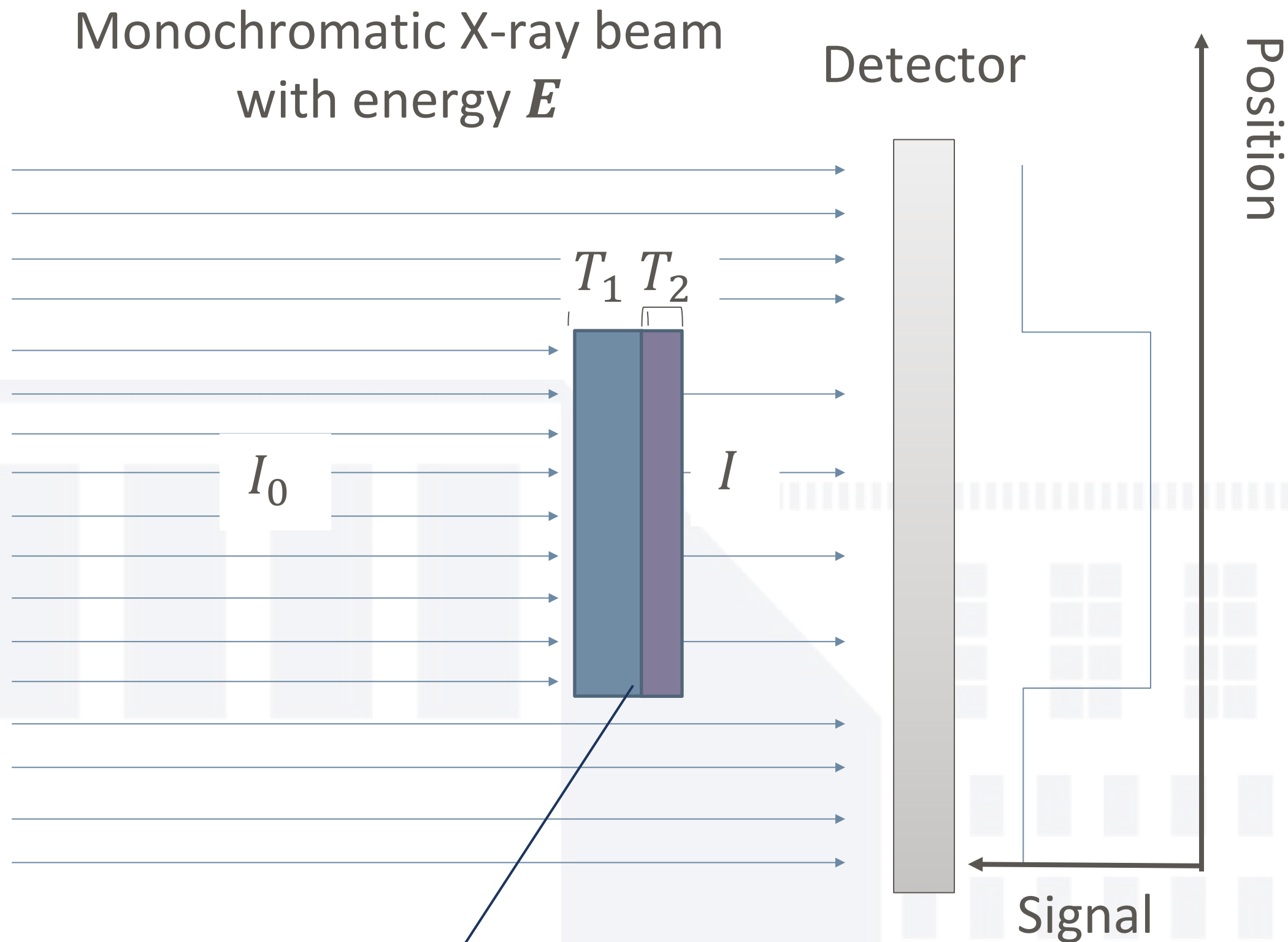
For a homogenous object of thickness  $T$

$$= I_0 e^{-\mu(E) T}$$

$$I = I_0 e^{-\frac{\mu}{\rho}(E) \rho T}$$

$\frac{\mu}{\rho}(E)$  → mass attenuation coefficient  
 independent on the physical state of the material/compound

# X-RAY SPECTRAL RADIOGRAPHY – BASICS (2)



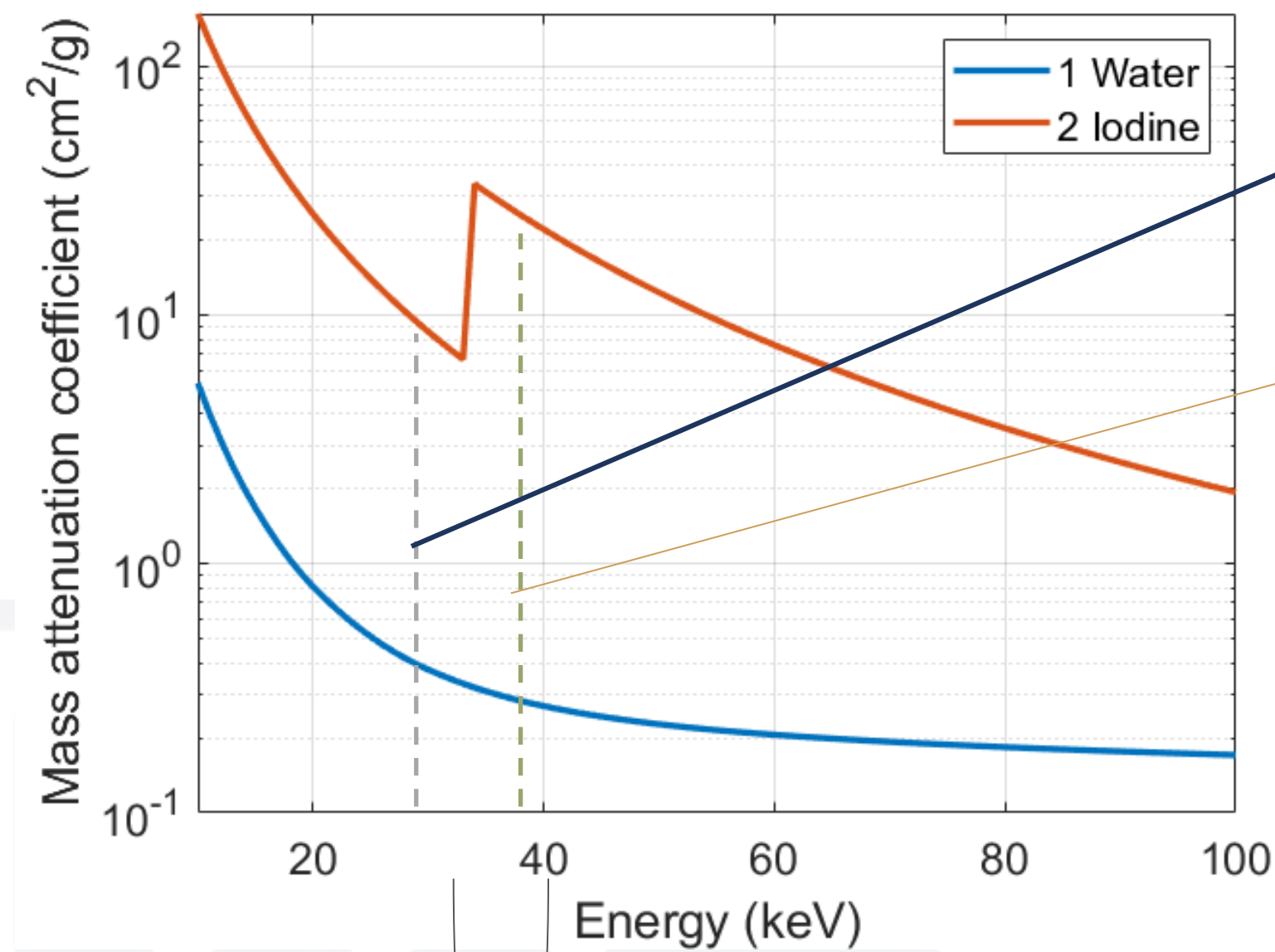
If we have 2 objects made of different materials...

$$I = I_0 e^{\left[ -\frac{\mu}{\rho}(E) \Big|_1 \rho_1 T_1 - \frac{\mu}{\rho}(E) \Big|_2 \rho_2 T_2 \right]}$$

$$P \equiv -\ln\left(\frac{I}{I_0}\right) = \left[ \frac{\mu}{\rho}(E) \Big|_1 \rho_1 T_1 + \frac{\mu}{\rho}(E) \Big|_2 \rho_2 T_2 \right]$$

From a single image  $I$ , one cannot uncouple/distinguish the 2 materials, i.e. **we cannot solve the equation for  $\rho_1 T_1$  and  $\rho_2 T_2$**

# BASIS MATERIAL DECOMPOSITION



Low energy image  $P^l = \frac{\mu}{\rho}\bigg|_1^l \rho_1 T_1 + \frac{\mu}{\rho}\bigg|_2^l \rho_2 T_2$

High energy image  $P^h = \frac{\mu}{\rho}\bigg|_1^h \rho_1 T_1 + \frac{\mu}{\rho}\bigg|_2^h \rho_2 T_2$

In a matrix form:

$$\begin{pmatrix} P^l \\ P^h \end{pmatrix} = \begin{pmatrix} \frac{\mu}{\rho}\bigg|_1^l & \frac{\mu}{\rho}\bigg|_2^l \\ \frac{\mu}{\rho}\bigg|_1^h & \frac{\mu}{\rho}\bigg|_2^h \end{pmatrix} \begin{pmatrix} \rho_1 T_1 \\ \rho_2 T_2 \end{pmatrix}$$

Two monochromatic energy channels (e.g., with synchrotron)

## References

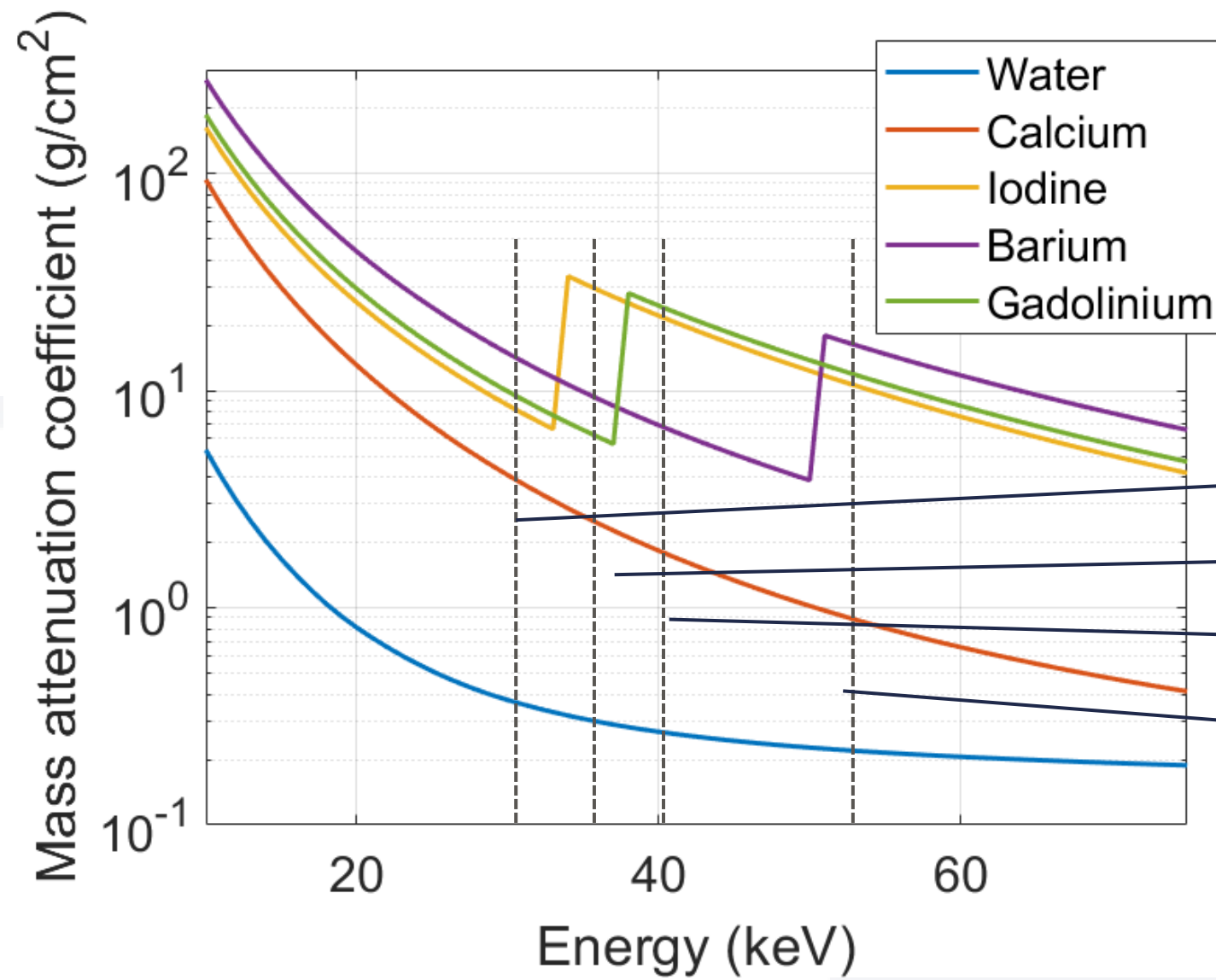
- [Alvarez, R E; Macovski, A \(1976\). Energy-selective reconstructions in X-ray computerised tomography. Physics in Medicine and Biology, 21\(5\), 733–744.](#)
- [Lehmann LA, Alvarez RE, Macovski A, Brody WR, Pelc NJ, Riederer SJ, Hall AL. Generalized image combinations in dual KVP digital radiography. Med Phys. 1981 Sep-Oct;8\(5\):659-67](#)

Matrix inversion:

$$\begin{pmatrix} \rho_1 T_1 \\ \rho_2 T_2 \end{pmatrix} = \begin{pmatrix} \frac{\mu}{\rho}\bigg|_1^l & \frac{\mu}{\rho}\bigg|_2^l \\ \frac{\mu}{\rho}\bigg|_1^h & \frac{\mu}{\rho}\bigg|_2^h \end{pmatrix}^{-1} \begin{pmatrix} P^l \\ P^h \end{pmatrix}$$

# MULTIPLE BASIS MATERIAL DECOMPOSITION

- The algorithm can be **extended to multiple energy channels and multiple decomposition materials**



$n$  energy channels

$$\begin{bmatrix} \mu_1 \\ \vdots \\ \mu_n \end{bmatrix} = \begin{bmatrix} \frac{\mu}{\rho} \Big|_1^1 & \dots & \frac{\mu}{\rho} \Big|_1^m \\ \vdots & \ddots & \vdots \\ \frac{\mu}{\rho} \Big|_n^1 & \dots & \frac{\mu}{\rho} \Big|_n^m \end{bmatrix} \begin{bmatrix} \rho^1 \\ \vdots \\ \rho^m \end{bmatrix}$$

$m(\leq n)$  materials

Decomposition matrix

$$\vec{\mu} = A\vec{\rho}$$

$$\vec{\rho} = A^{-1}\vec{\mu}$$

Inversion performed through least-squares

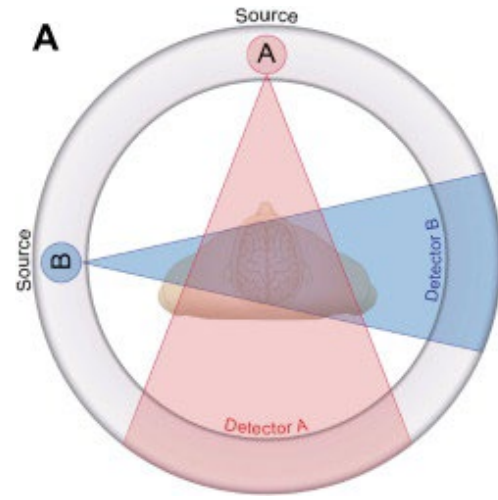
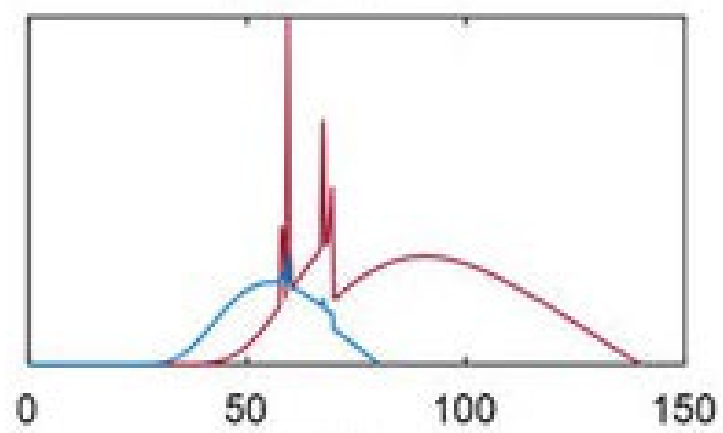
NOTE:

The choice of basis materials is crucial for the decomposition quality!  
E.g., if materials are too similar the noise will be amplified

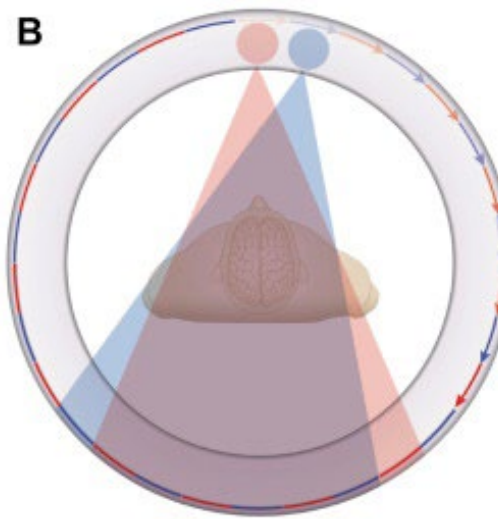
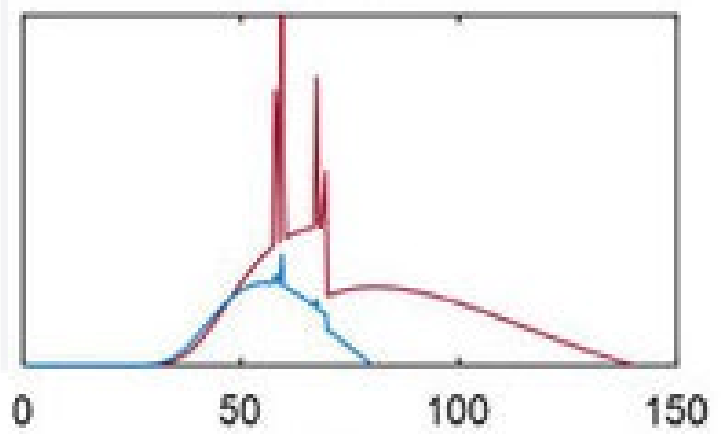
# SPECTRAL IMAGING SYSTEMS

## X-ray spectrum-based

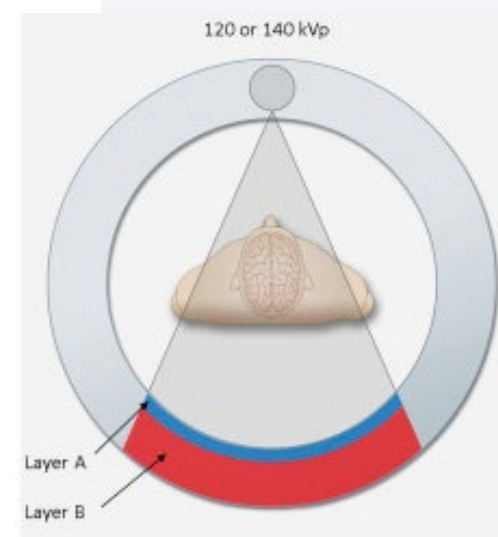
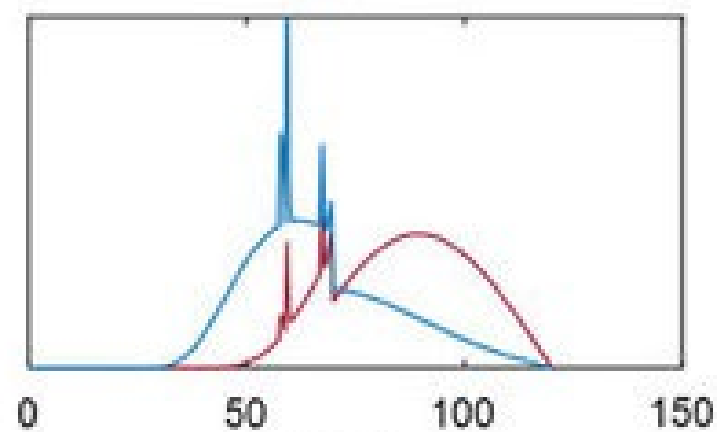
- 2 X-ray tubes with different voltages



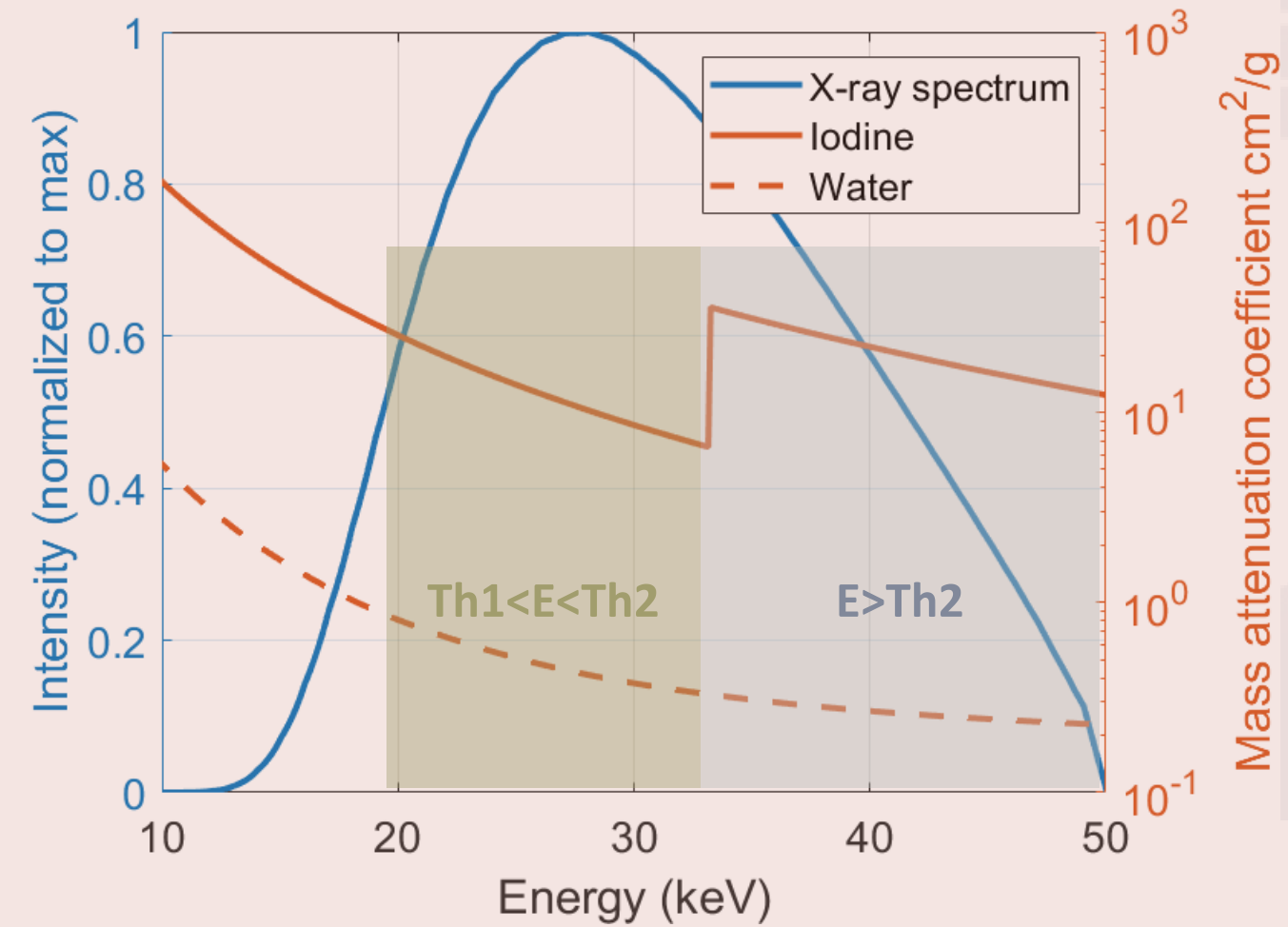
- Voltage switching



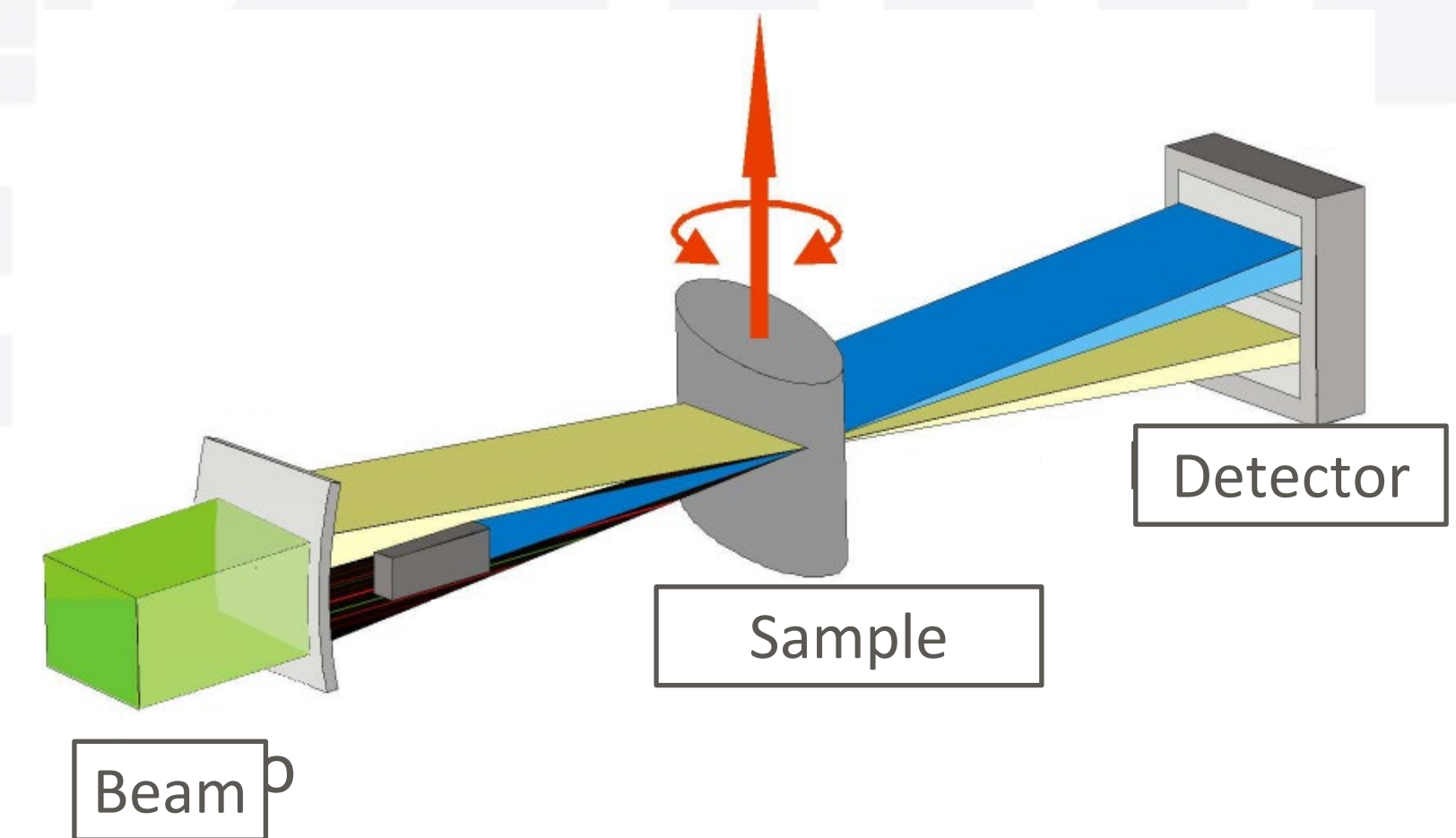
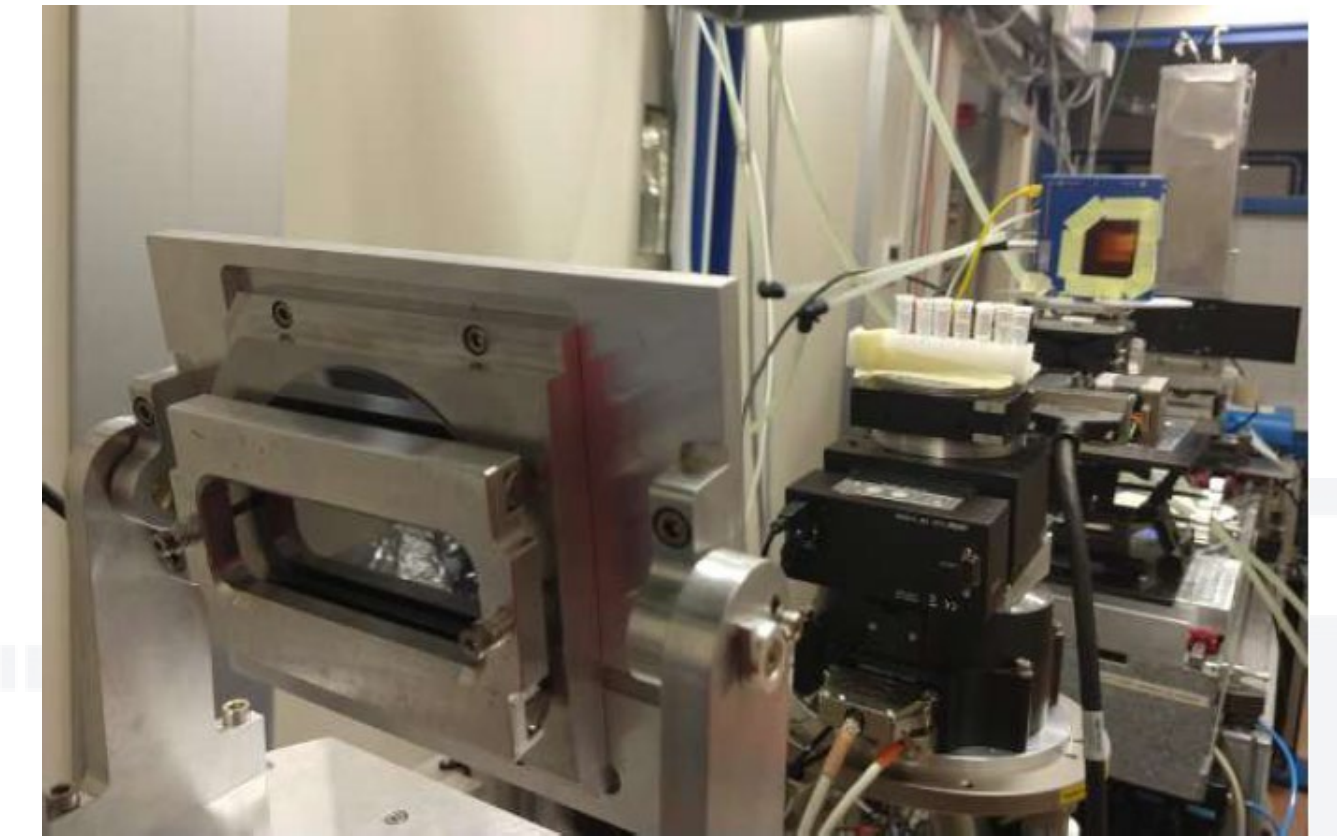
- Dual layer detectors



## Detector-based pixel-by-pixel energy discrimination



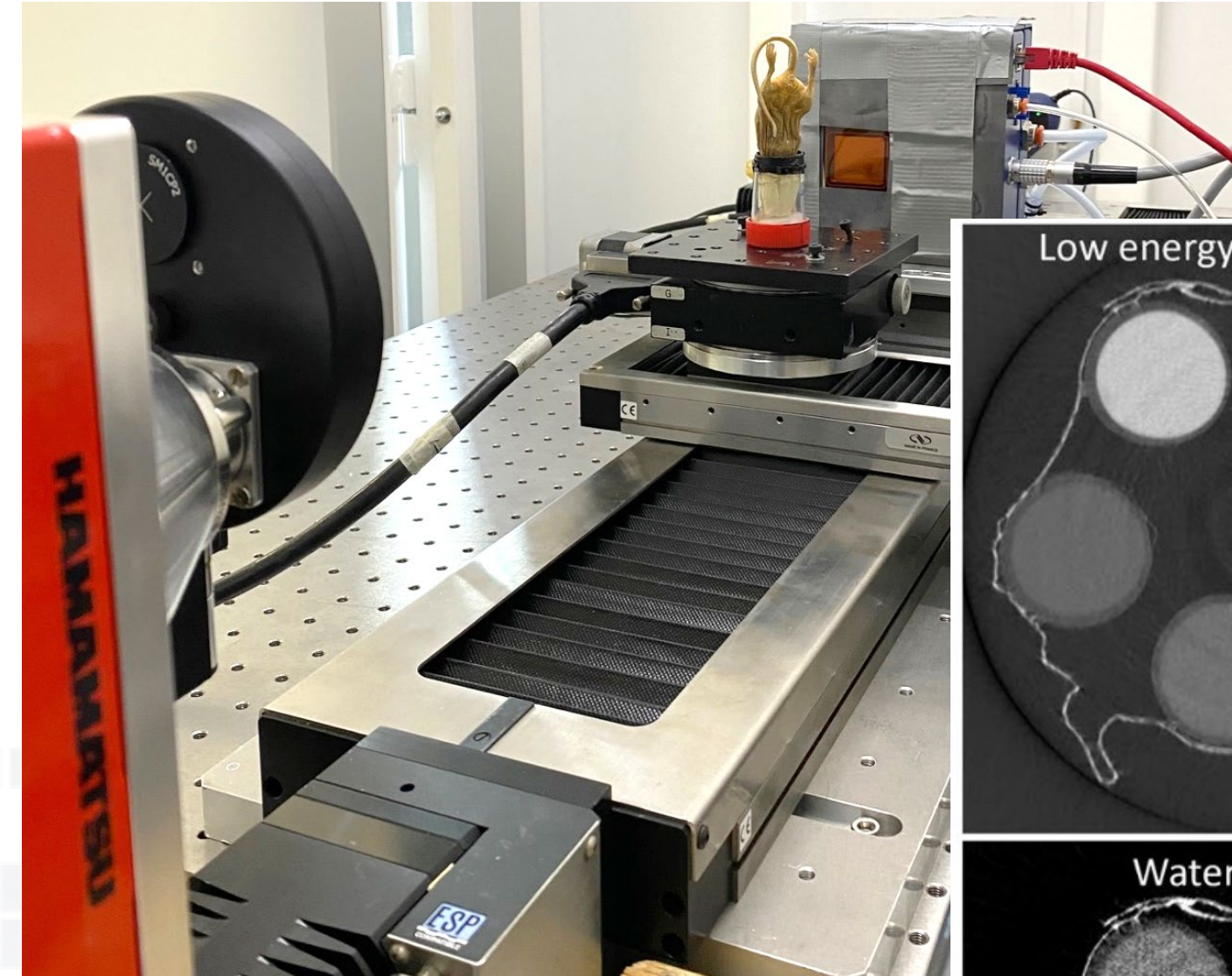
## Crystal-based energy dispersive systems based on bent crystals (@ synchrotrons)



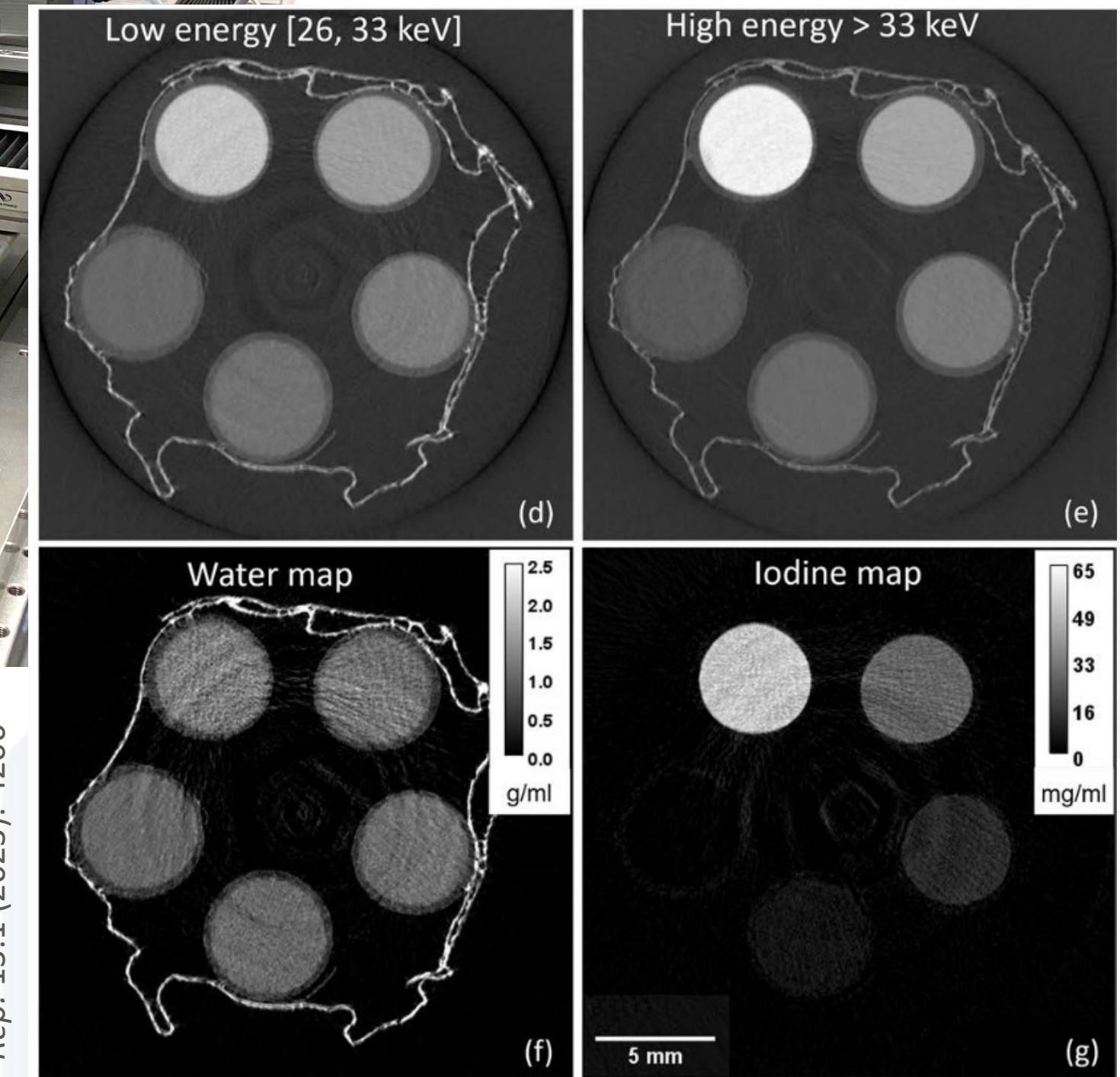
# SPECTRAL DETECTORS

- Spectral detectors can acquire multiple **images over different energy channels in a single shot**
- **High-Z sensors** (CdTe, CZT, GaAs, ...) are used for spectral imaging due to their high efficiency at high energies (>30 keV)

CHIP/PRODUCER	PIXEL SIZE ( $\mu\text{m}$ )	NUMBER OF THRESHOLDS
MEDIPIX3	55	2 (8 in 2x2 Binning)
PIXIRAD – PIXIEIII	62	2
DIRECT CONVERSION	100	2
DECTRIS - EIGER 2	75	2
TIMEPIX4	55	Full Spectrum, ~1.5 KeV resolution
...		



F Brun, et al. *Phys. Med. & Biol.* 65.5 (2020): 055016. ← Funded by INFN



Brombal, Luca, et al. *Sci. Rep.* 13.1 (2023): 4206

# SPECTRAL DETECTORS: HOW DO WE MEASURE X-RAY ENERGY?

## Energy measurement principle

Time-over-threshold

Example: *Timepix*

Photon counting with multiple thresholds

Example: *Medipix, Pixirad*

Charge integrating single photon sensitivity

Example: *CITIUS by REIKEN*

# ENERGY MEASUREMENT VIA TIME-OVER-THRESHOLD (TOT)

The signal amplitude and duration is proportional to the energy released in the sensor from the X-ray



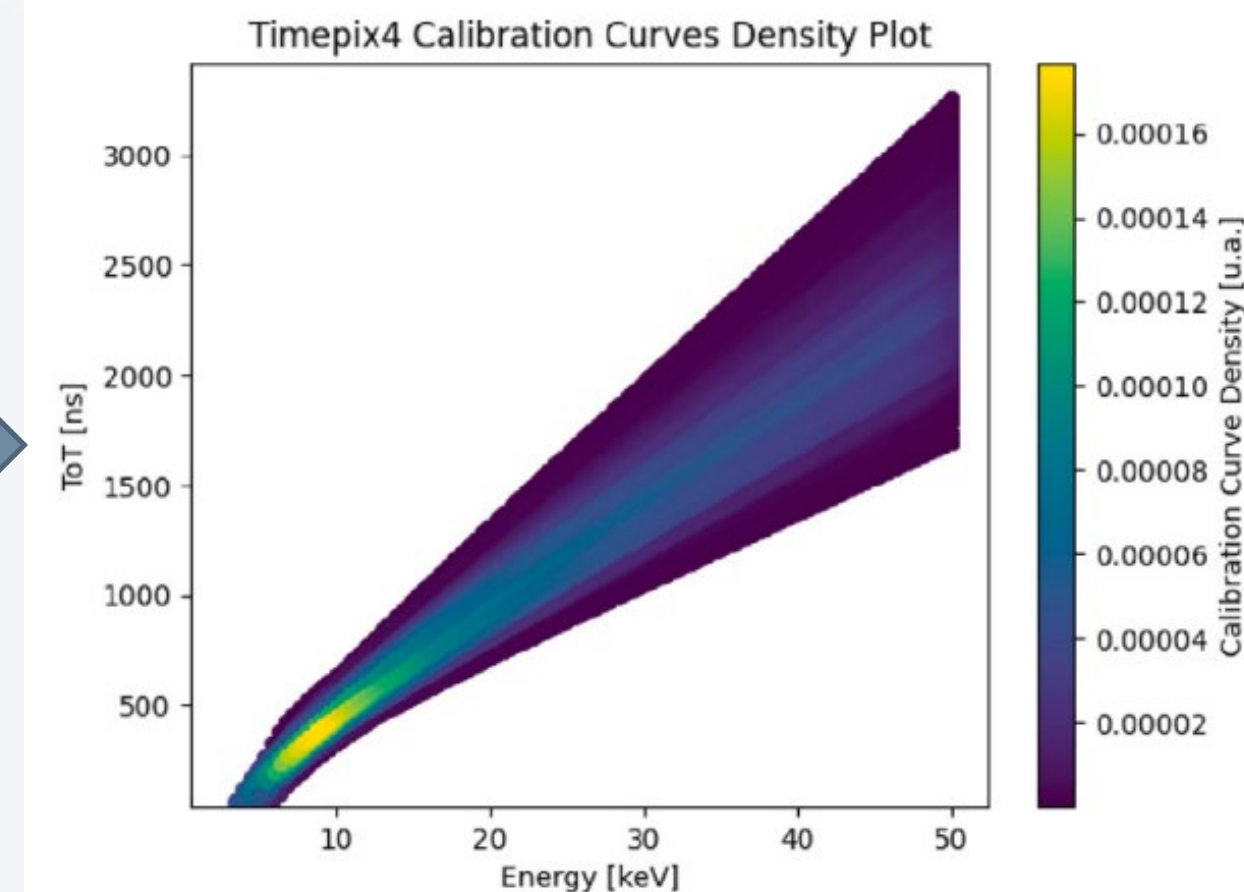
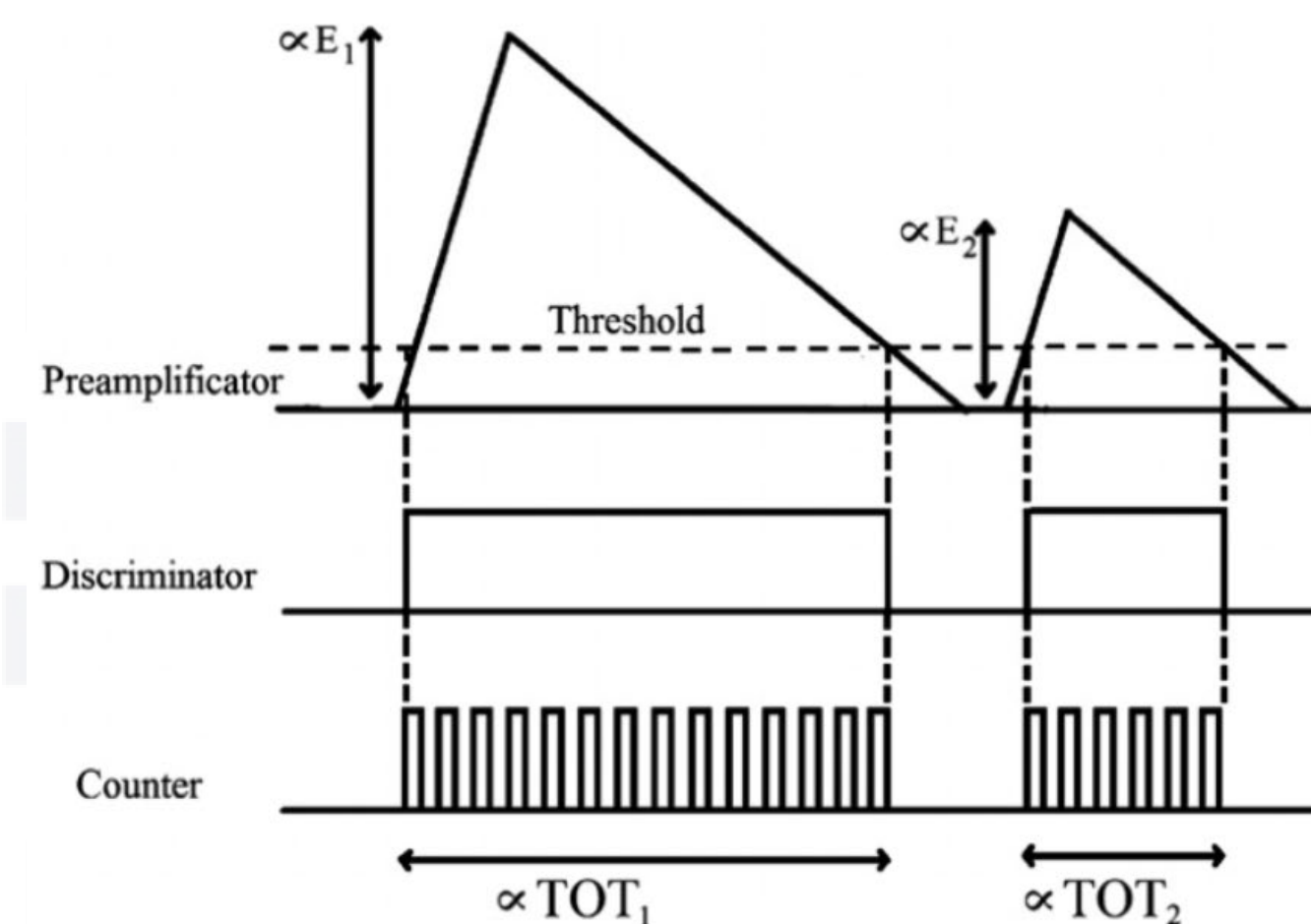
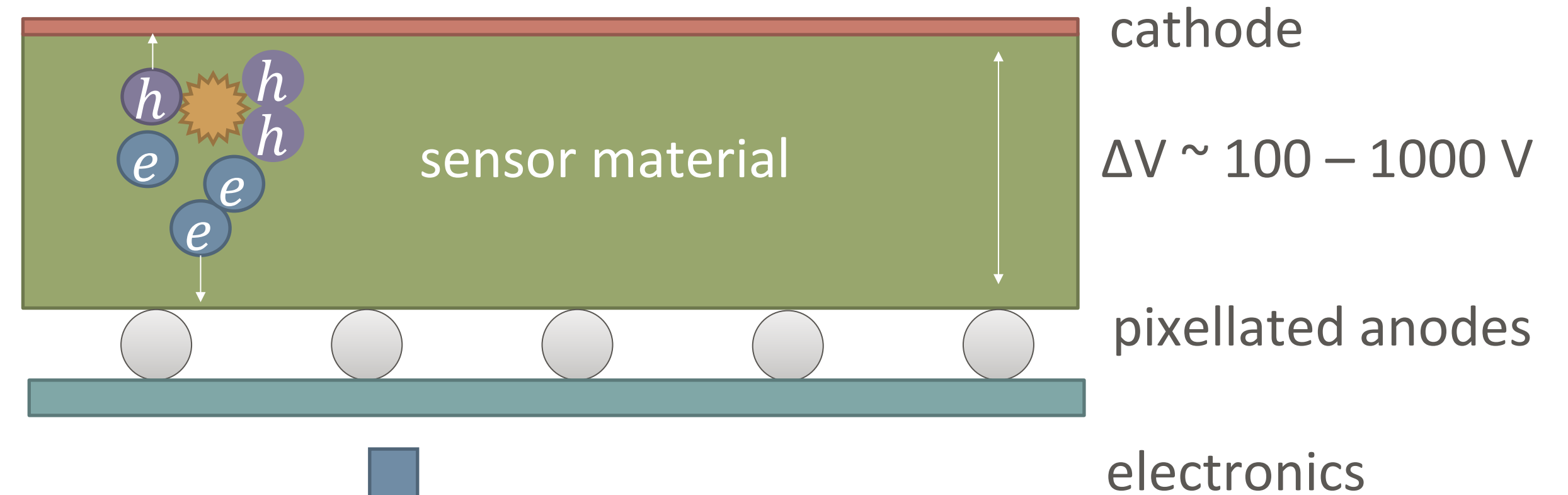
When the signal amplitude exceeds a threshold, a clock starts. It stops when the signal is below the threshold



The number of clock cycles is proportional to the amplitude i.e. the energy



For each event you have the full energy information (*hyperspectral performance*)



Delogu, P., et al. *Nuclear Instruments and Methods in Physics Research Section A: Accelerators, Spectrometers, Detectors and Associated Equipment* 1068 (2024): 169716.



# ENERGY MEASUREMENT VIA DISCRIMINATION (PHOTON-COUNTING)

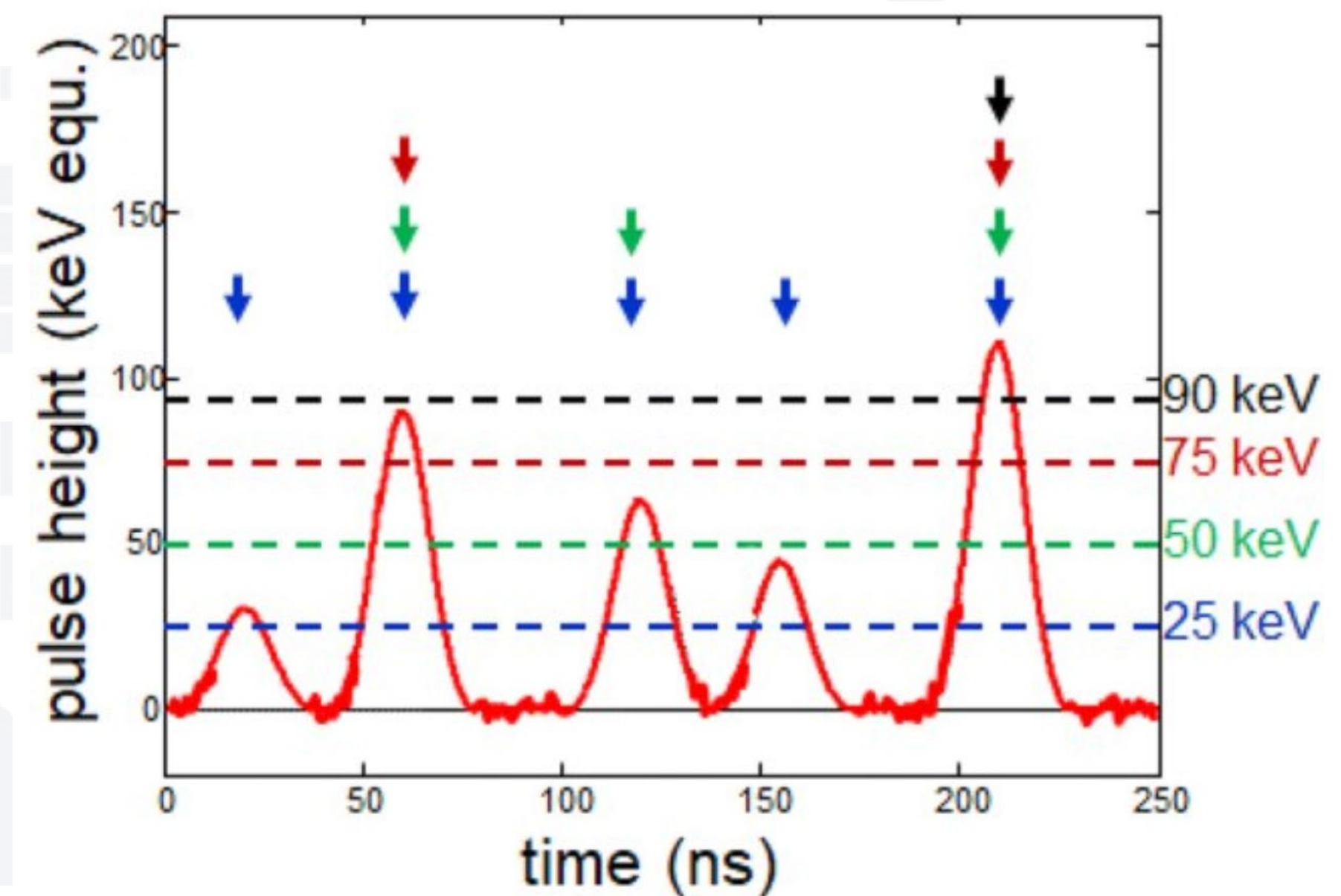
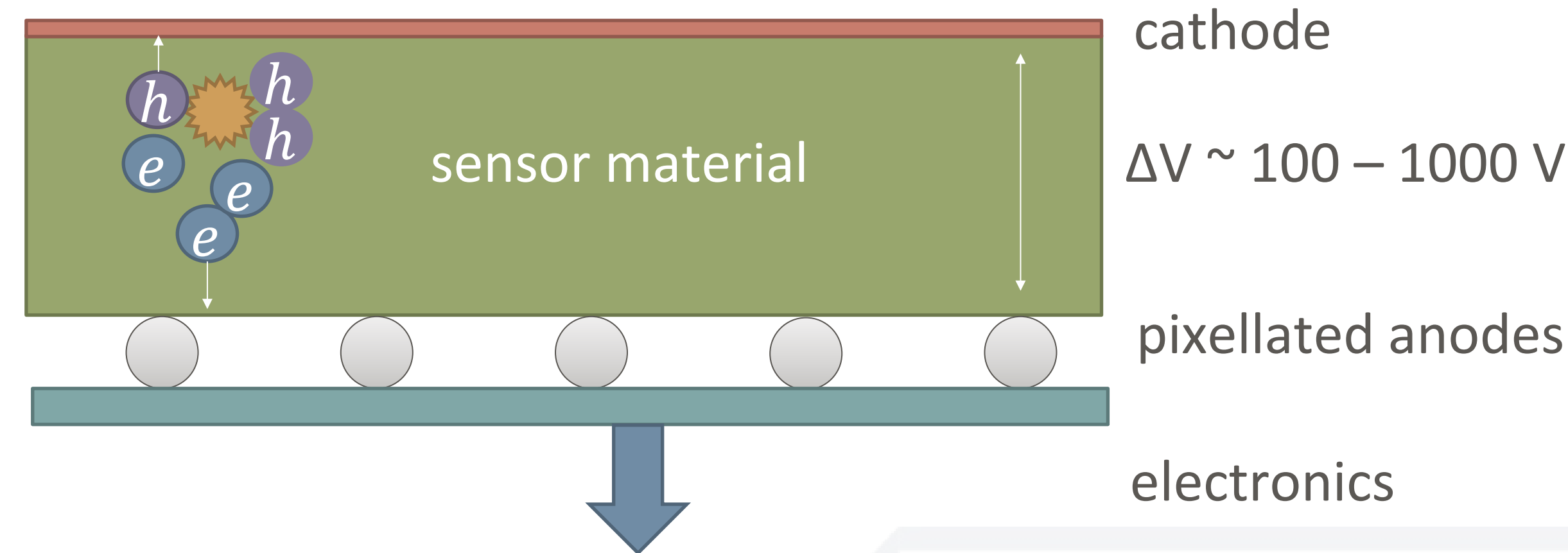
The signal amplitude and duration is proportional to the energy released in the sensor from the X-ray



When the signal amplitude exceeds a programmable energy-calibrated threshold a counter is incremented (+1)

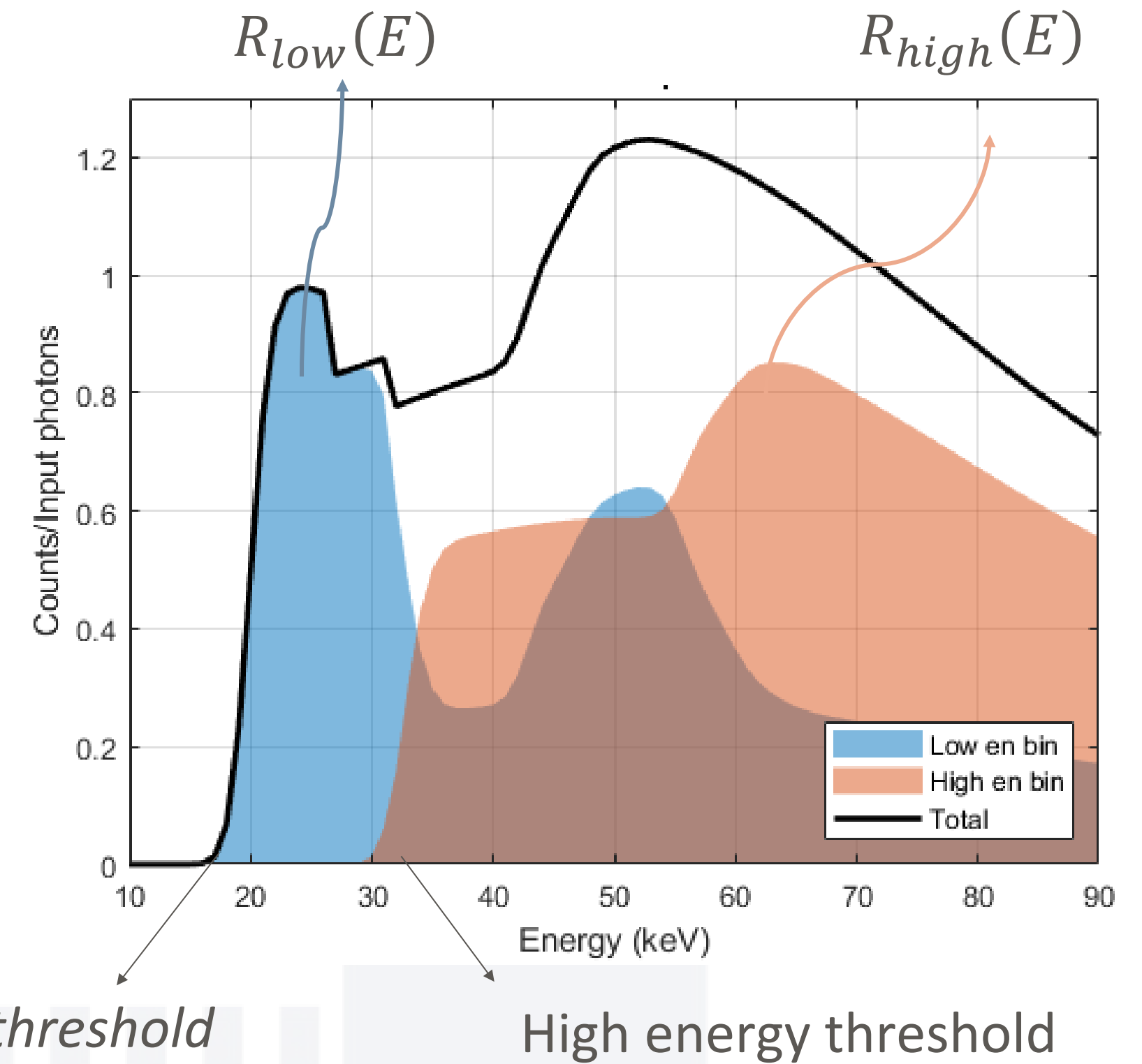
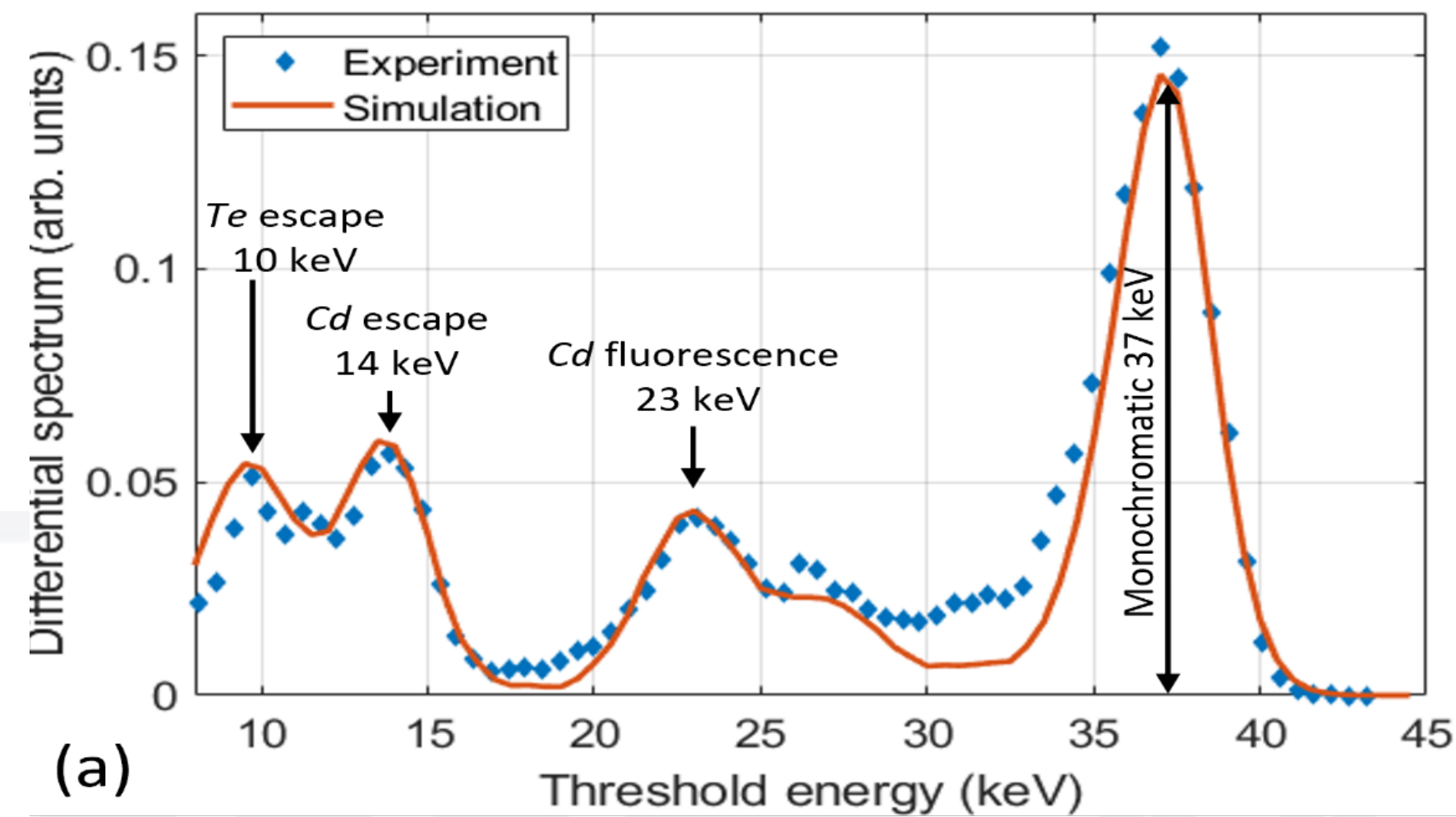


By having multiple thresholds per pixel multiple energy bins are acquired



❖ Flohr, Thomas, et al. "Basic principles and clinical potential of photon-counting detector CT." *Chinese Journal of Academic Radiology* 3.1 (2020): 19-34.

# MODELLING DETECTOR'S ENERGY RESPONSE



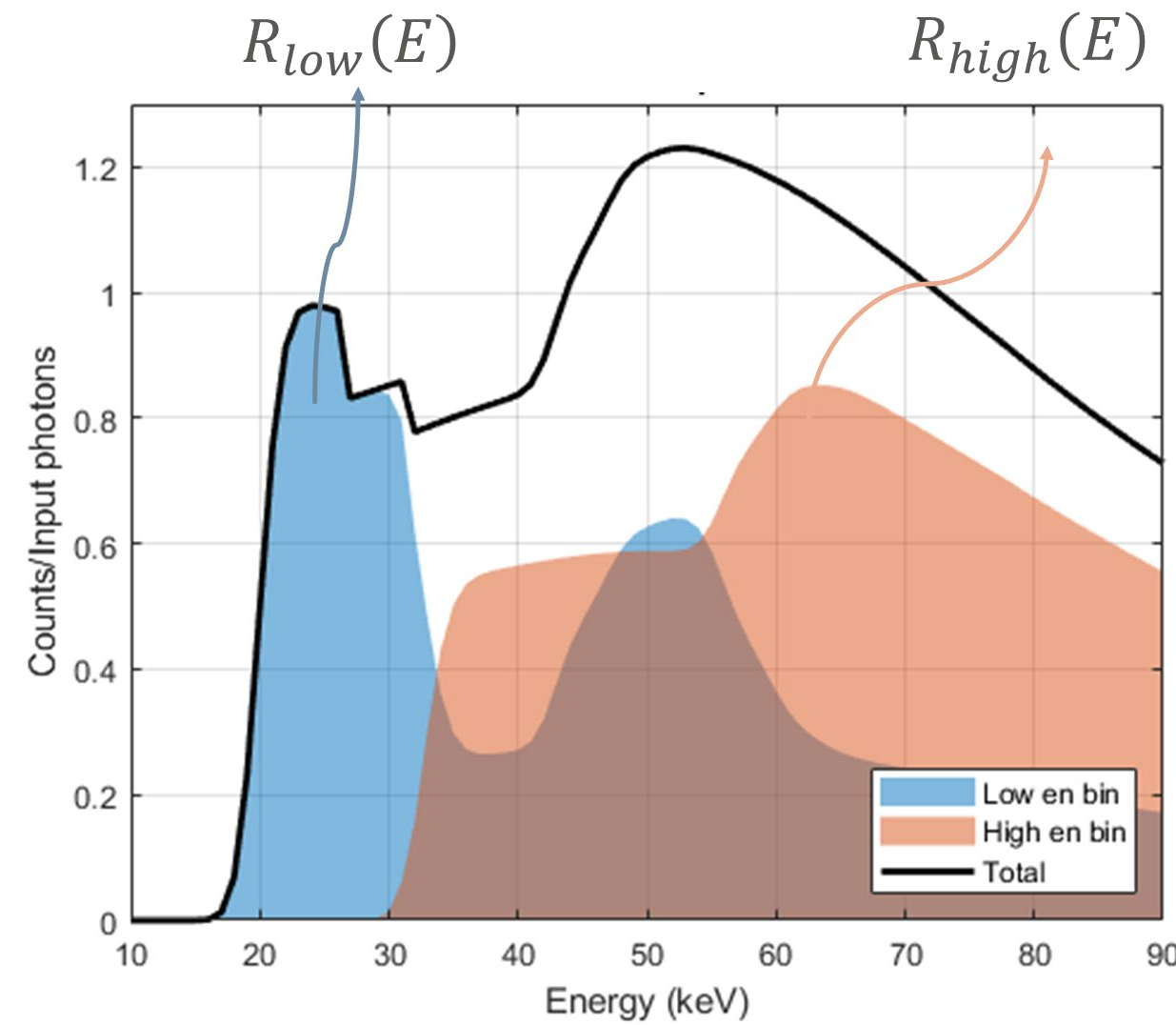
1. Measure the energy response of the detector to monochromatic radiation
2. Model its energy response at arbitrary energy levels



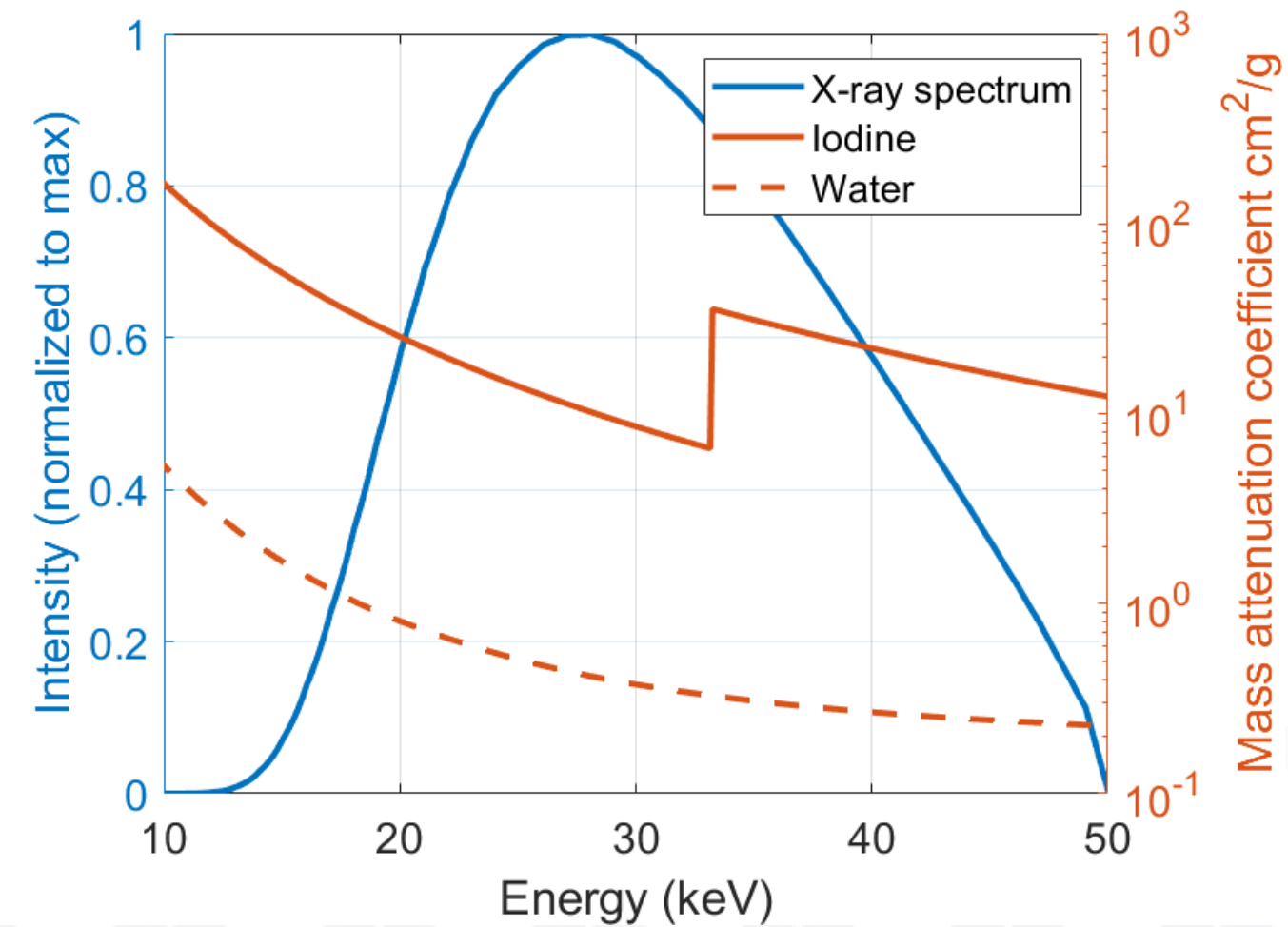
3. For given threshold setting, compute the probability that a photon of energy  $E$  is counted in a given bin

# COMPUTING THE BASIS-DECOMPOSITION MATRIX

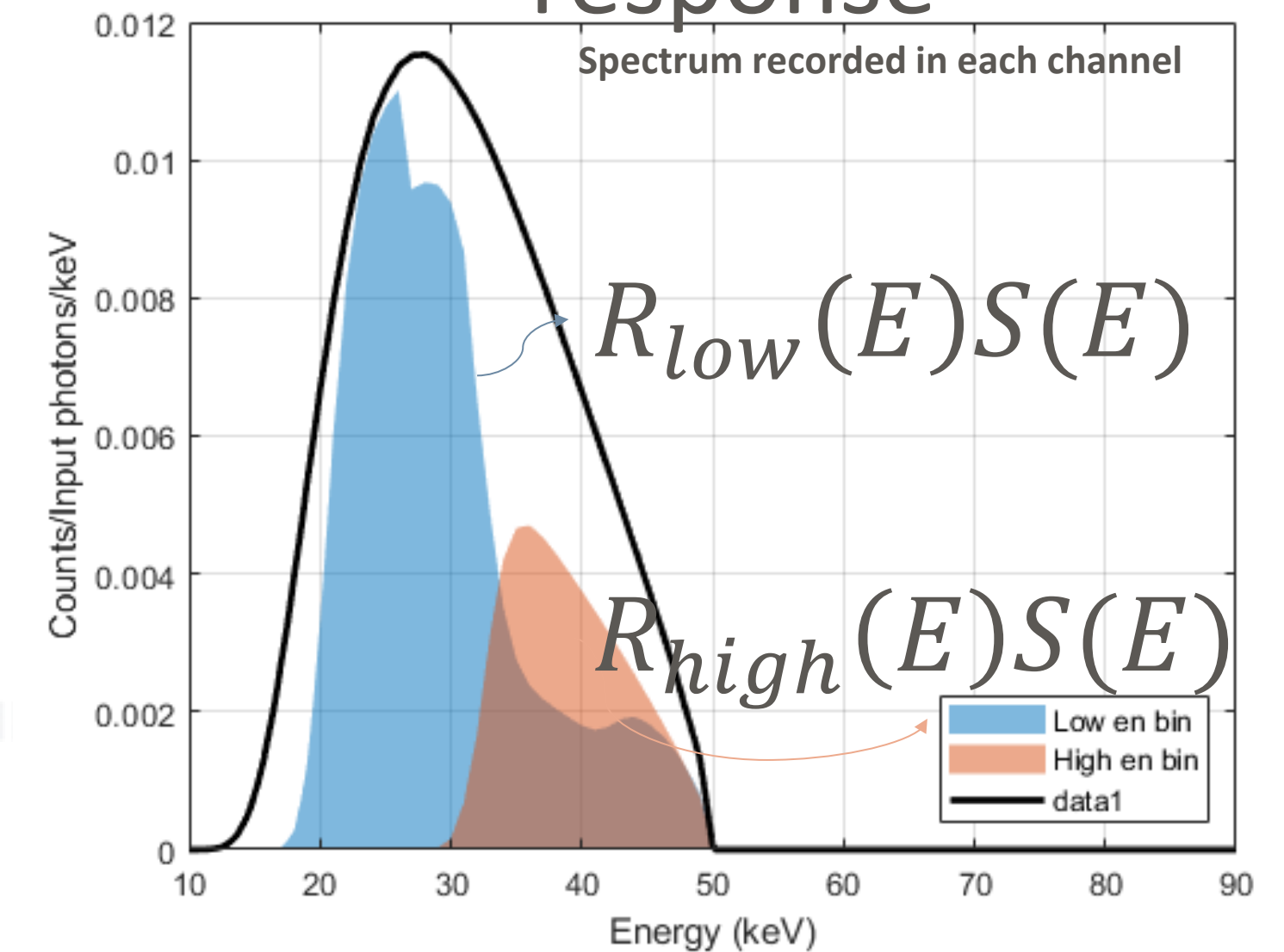
## Detector response



## X-ray spectrum



## System's spectral response

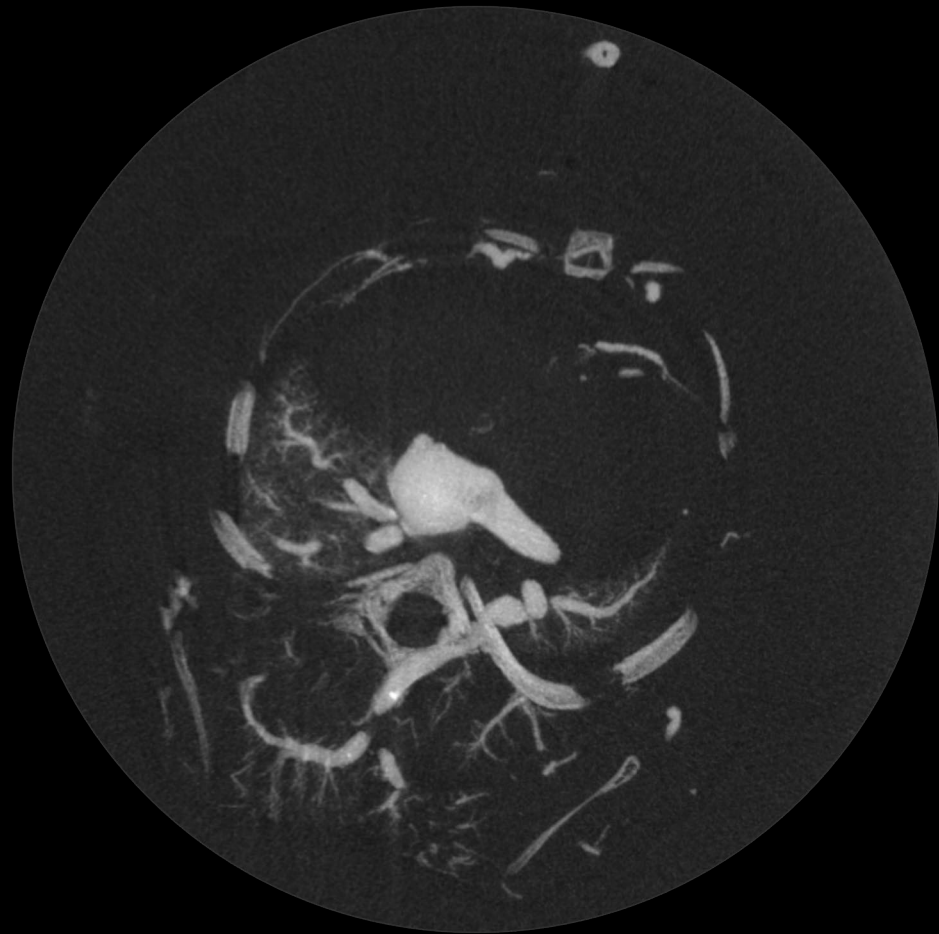


- **Decomposition matrix** calculated as a **weighted sum of the mass attenuation coefficients**.
- The **weights** are given by the **system's spectral response**

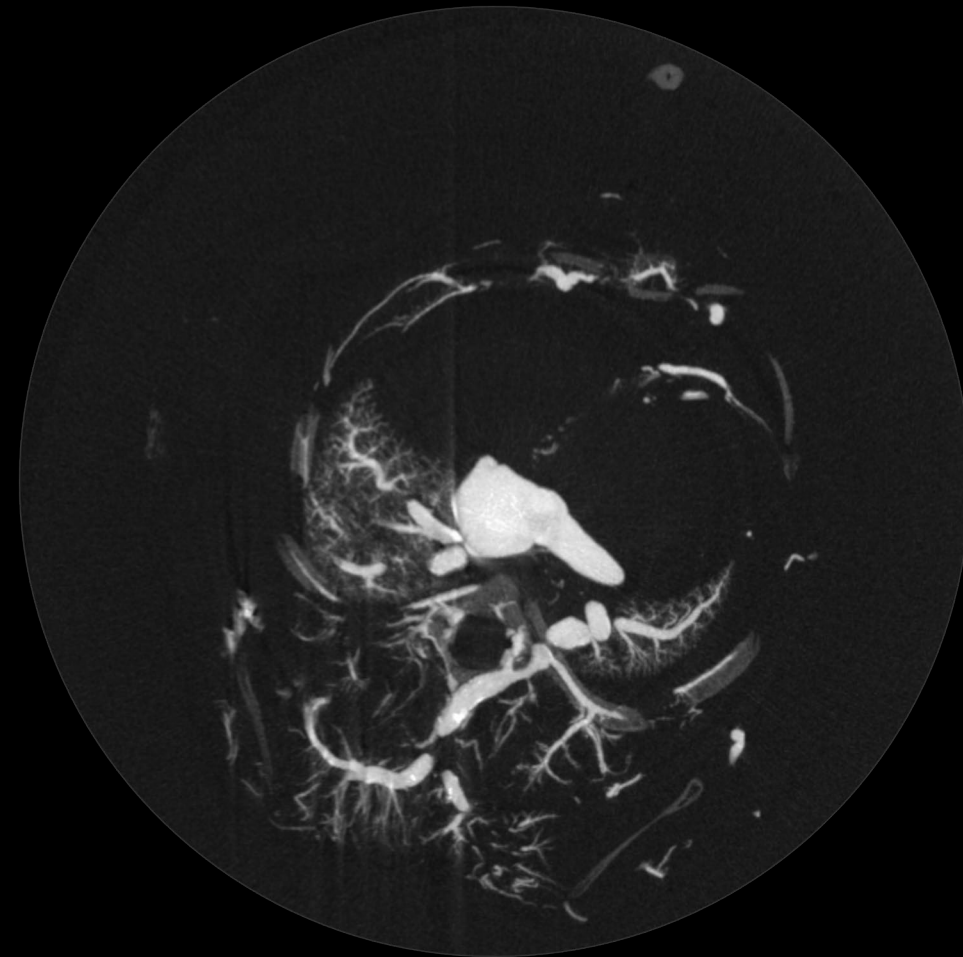
$$\left\langle \frac{\rho_j}{\rho_i} \right\rangle = \frac{\int S(E)R_i(E)\mu/\rho_j(E)dE}{\int S(E)R_i(E)dE}$$

# APPLICATIONS: CADIOVASCULAR IMAGING

Bin1 [27, 33] keV

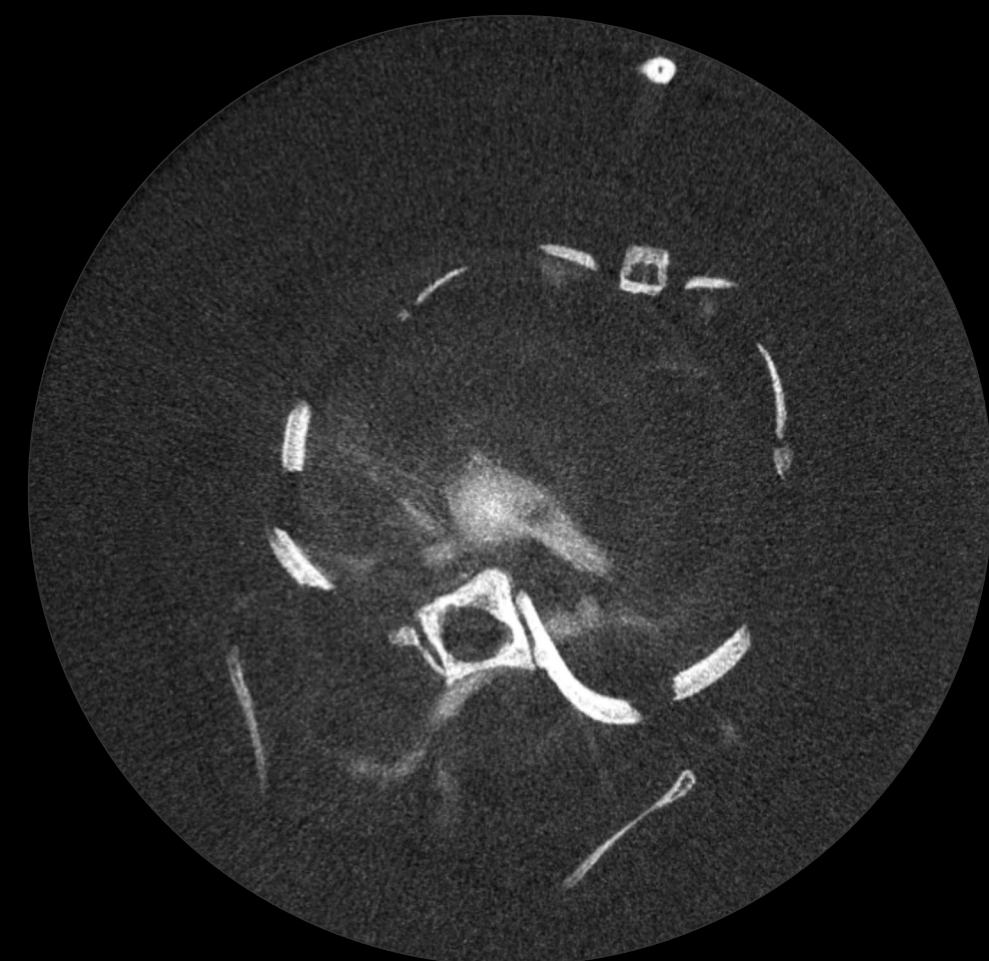


Bin2 >33 keV

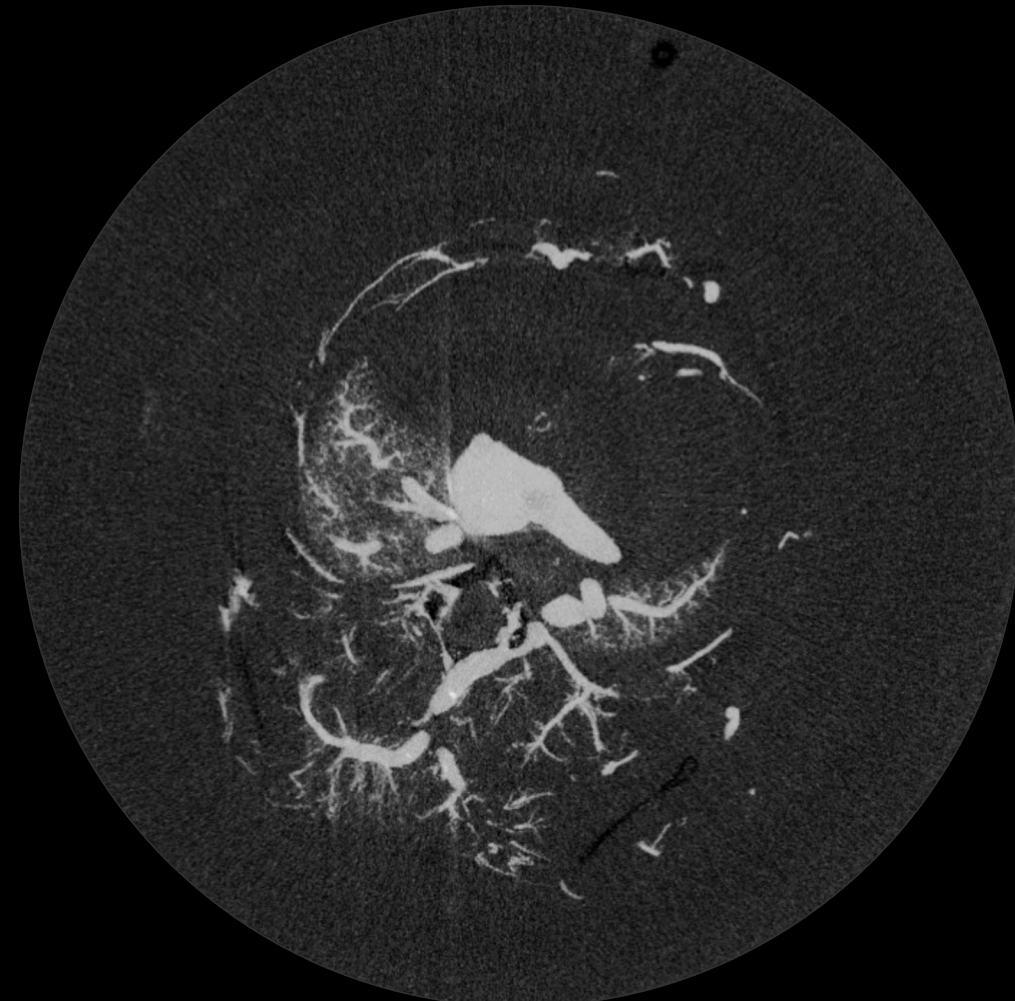


1 cm

Bone map



Iodine map



- 35  $\mu\text{m}$  voxel size
- 790 x 790 x 2500 voxel x 2 energies
- 12 Gb dataset

Ex-vivo murine model perfused with  $\mu\text{Angiofil}^{\text{®}}$



Thanks to Paola Perion

In collaboration with

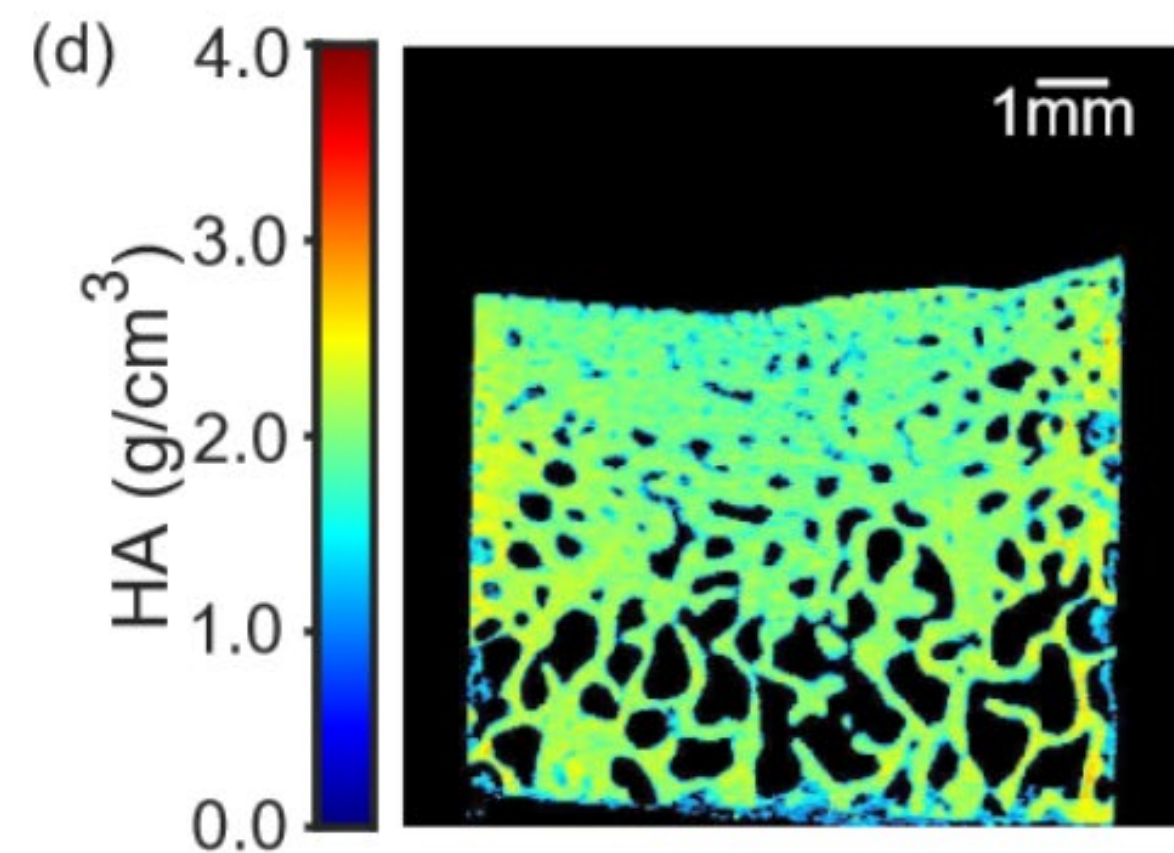
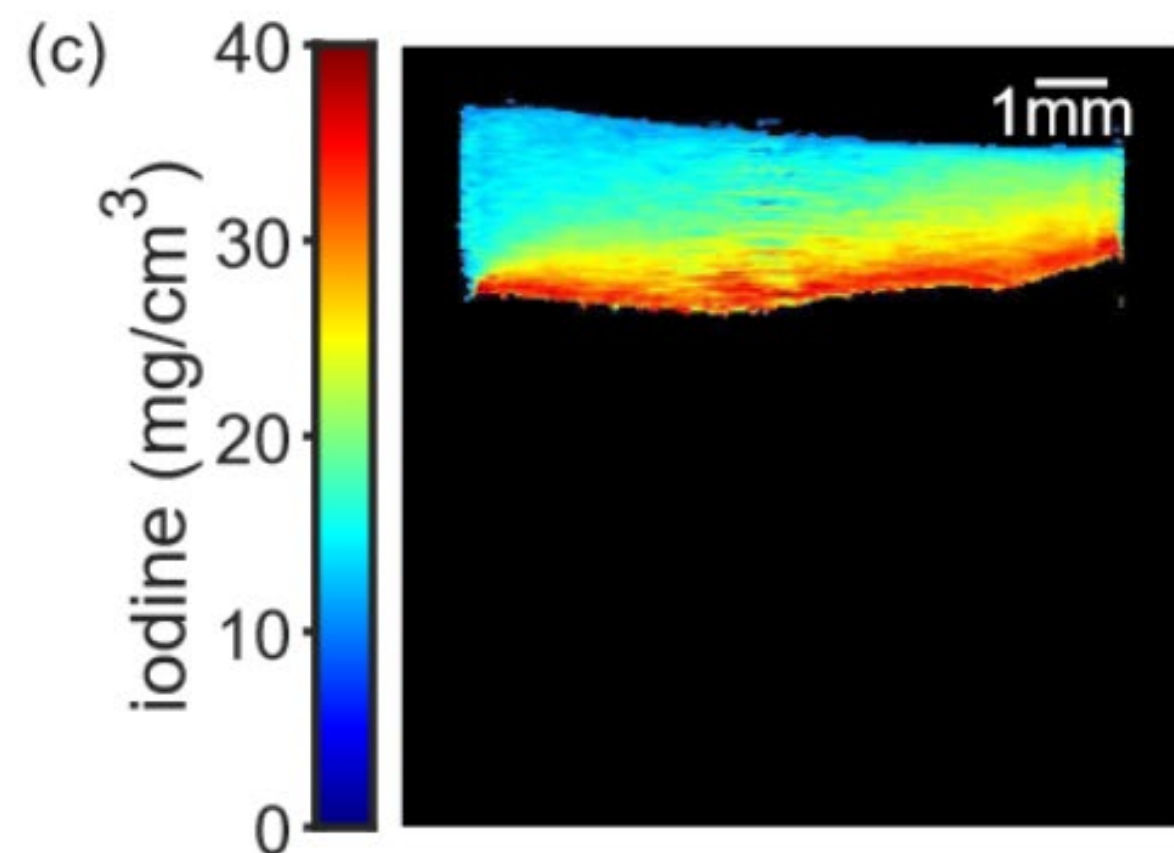
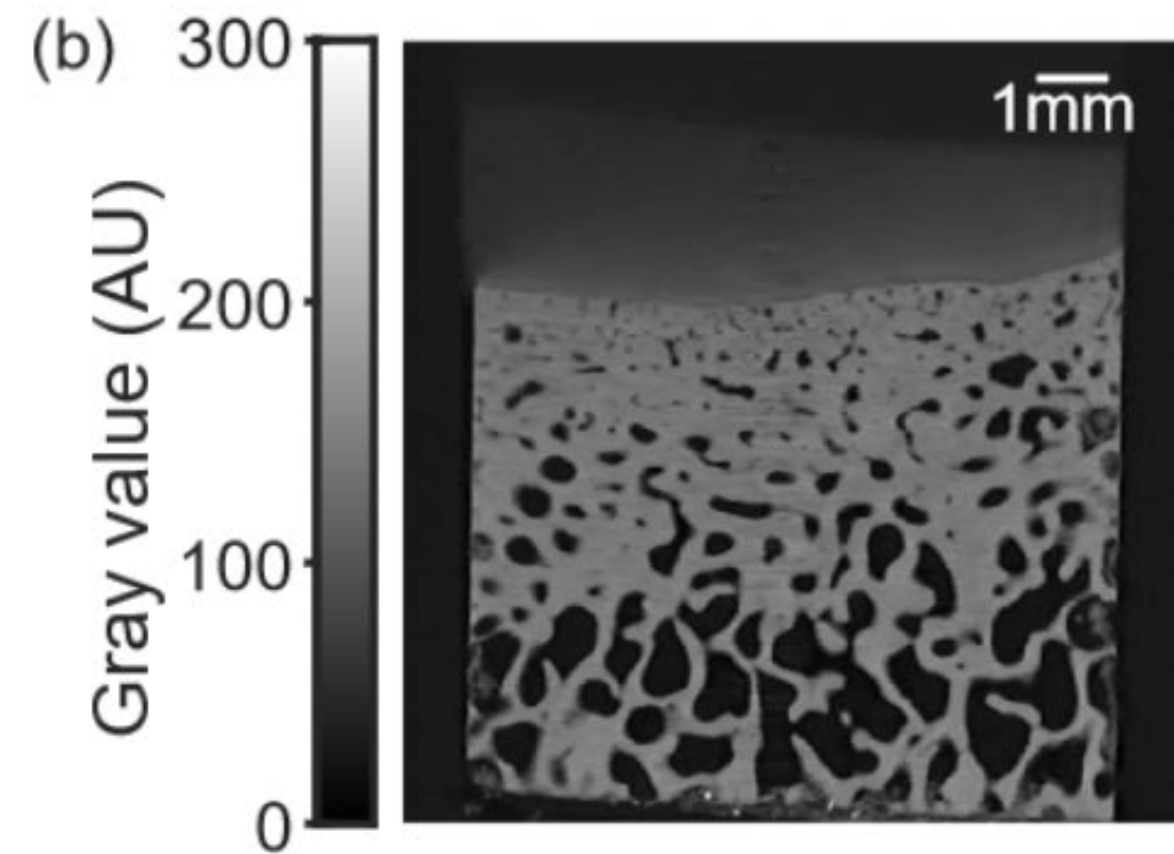
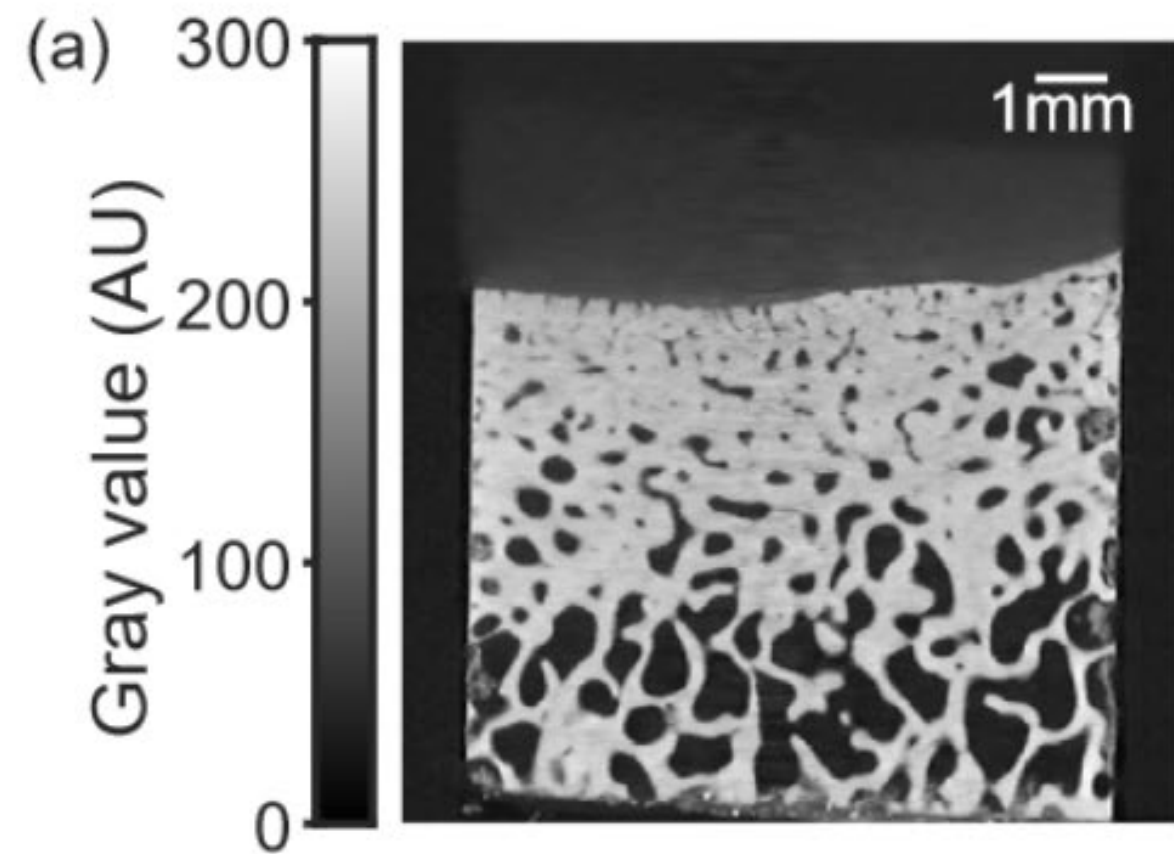
**HELMHOLTZ  
MUNICH**

# APPLICATION TO CA4+ LABELED OSTEOARTICULAR SAMPLES

voxel size 34  $\mu\text{m}$ , scan time  $\sim 120$  min

Low energy

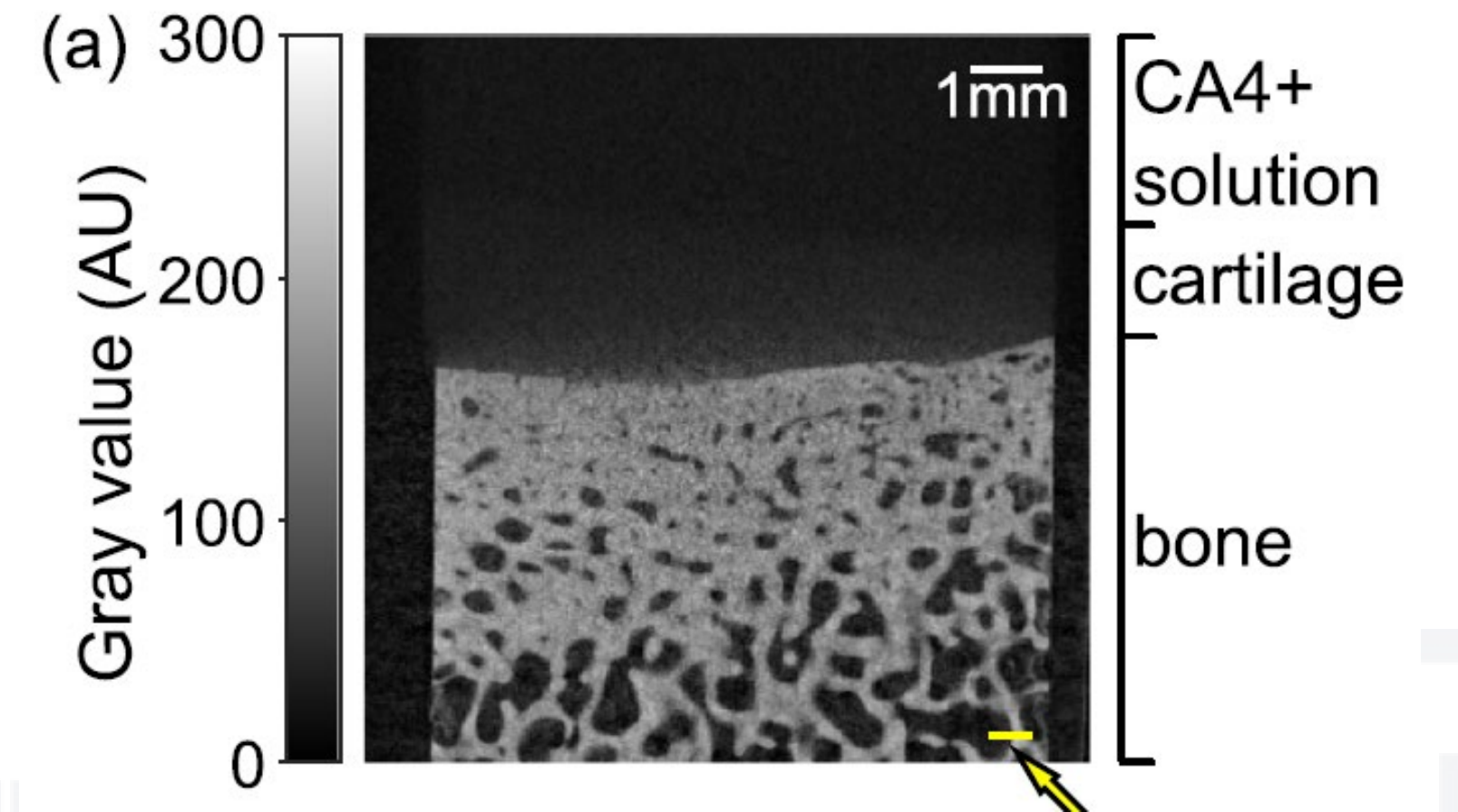
High energy



Iodine/cartilage

Hydroxyapatite/bone

Conventional  $\mu\text{CT}$



Bovine stifle joint sample labeled with a cationic iodinated contrast medium (CA4+)



In collaboration with:

SERVIZIO SANITARIO REGIONALE  
EMILIA - ROMAGNA  
Istituto Ortopedico Rizzoli di Bologna  
Istituto di Ricovero e Cura a Carattere Scientifico

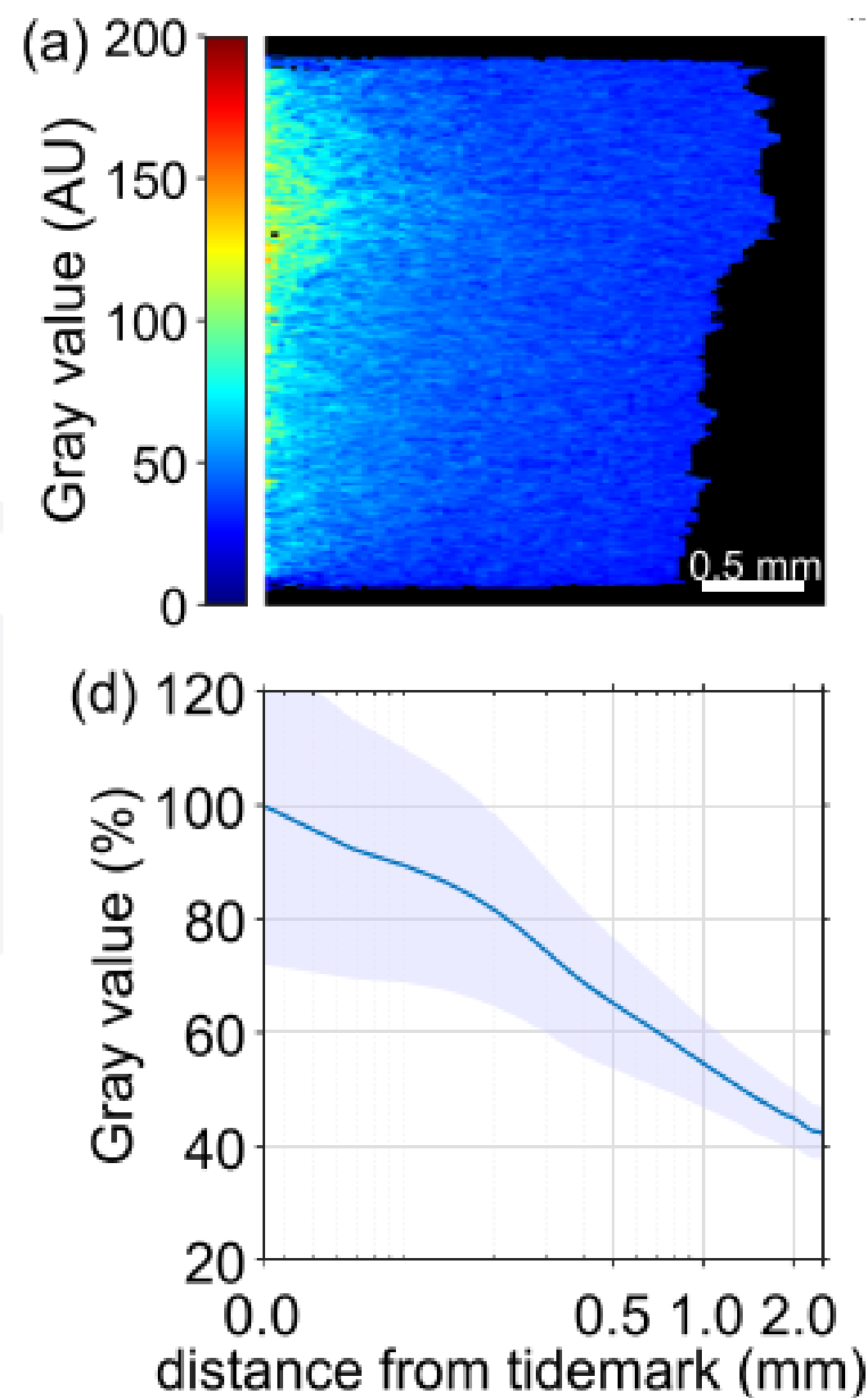


Fantoni, Simone, et al. *The European Physical Journal Plus* 139.8 (2024): 1-10.

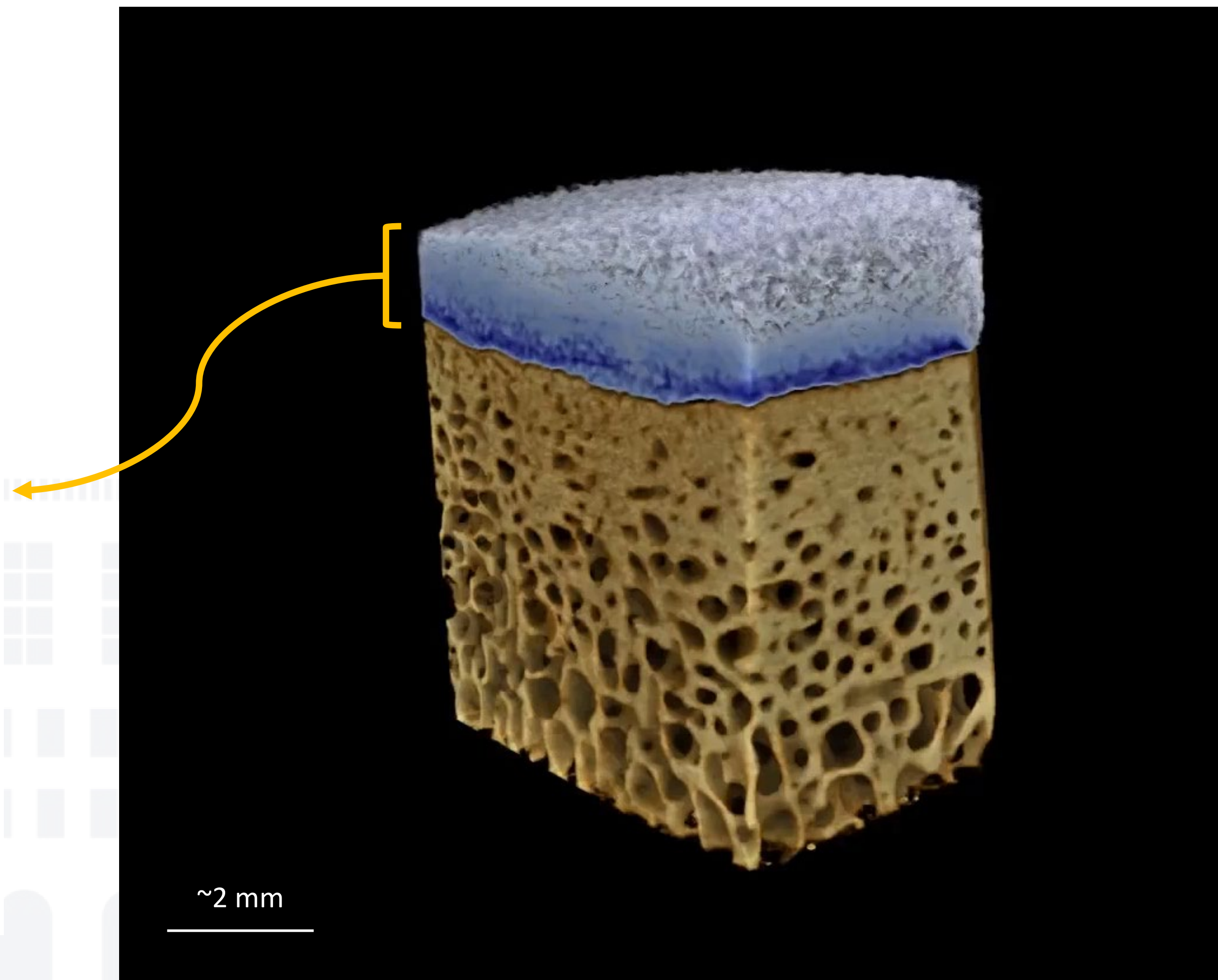
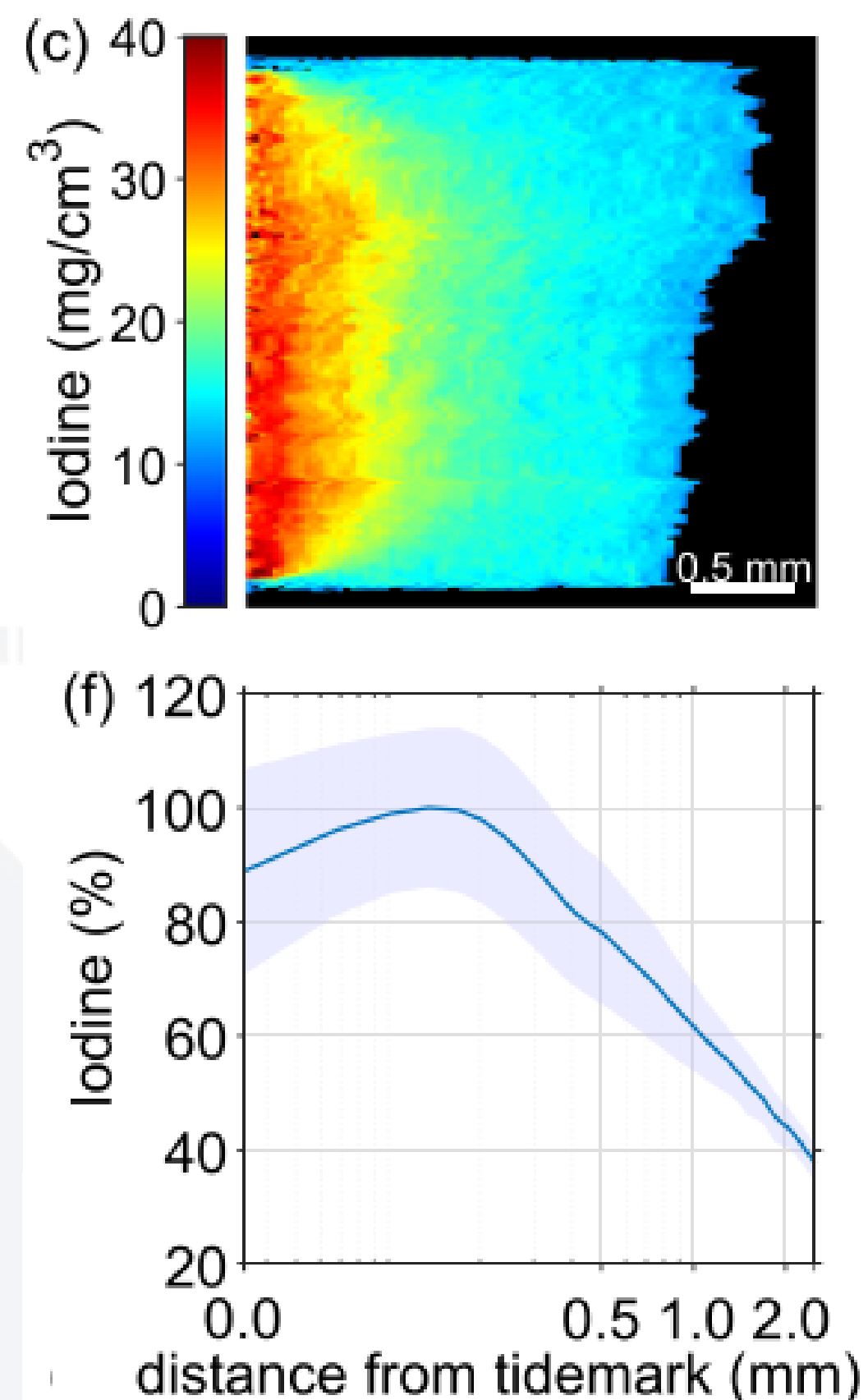
# APPLICATION TO CA4+ LABELED OSTEOARTICULAR SAMPLES

voxel size 34  $\mu\text{m}$ , scan time  $\sim 120$  min

## CONVENTIONAL



## SPECTRAL



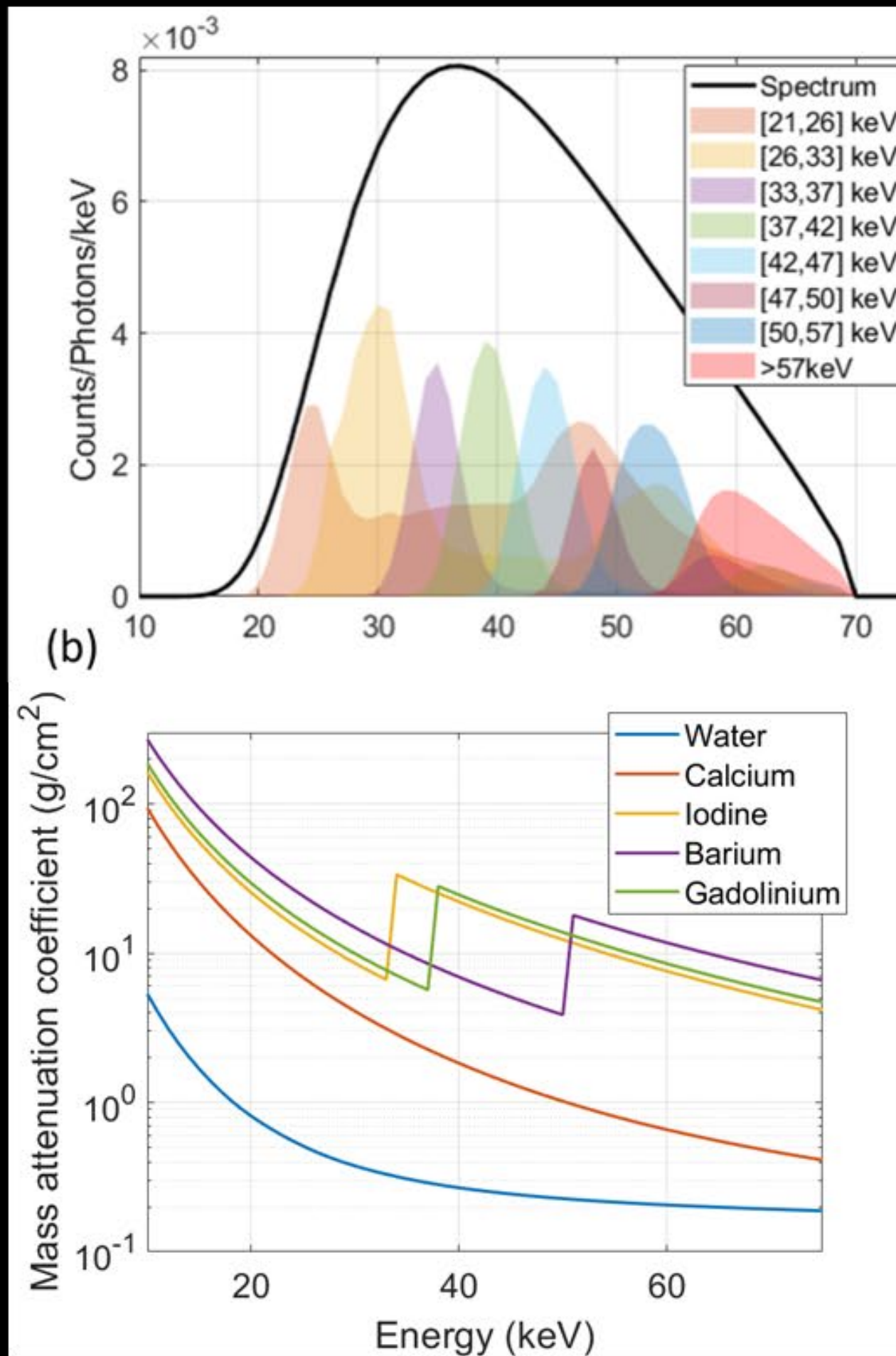
Fantoni et al. "Quantitative spectral micro-CT of a CA4+ loaded osteochondral sample with a tabletop system". Accepted in EPJ plus (2024)

In collaboration with:

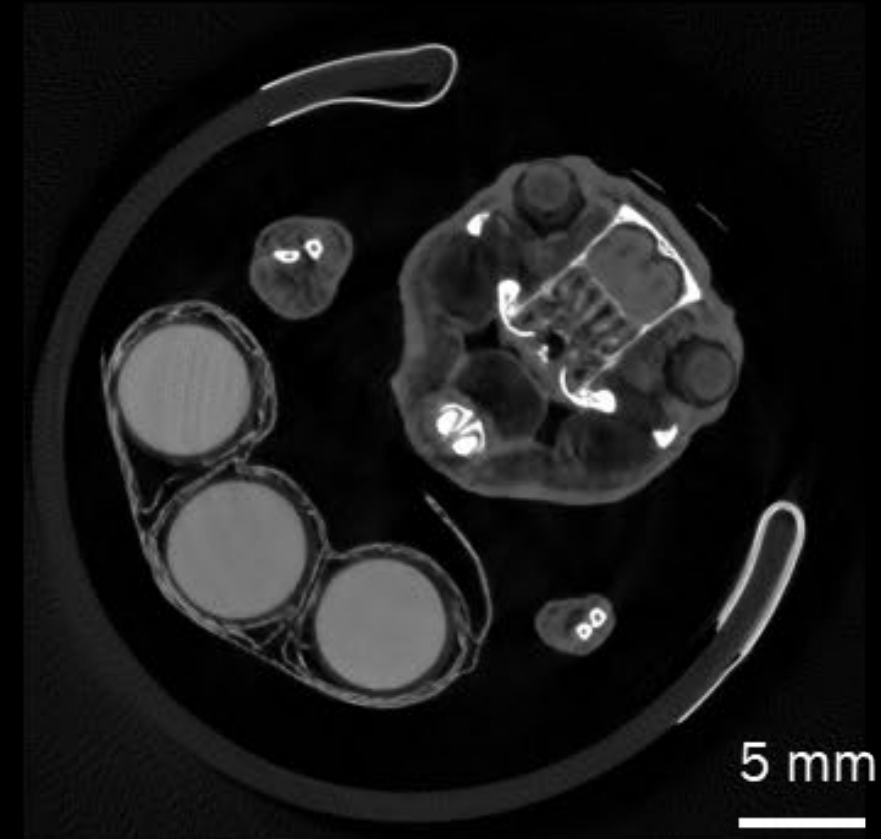
SERVIZIO SANITARIO REGIONALE  
EMILIA - ROMAGNA  
Istituto Ortopedico Rizzoli di Bologna  
Istituto di Ricovero e Cura a Carattere Scientifico



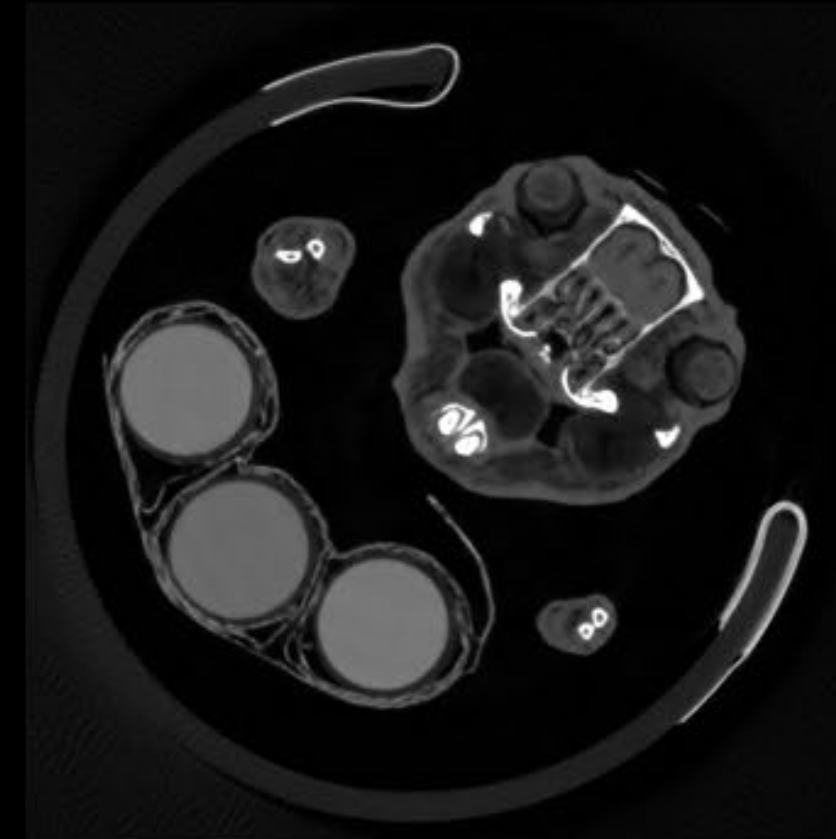
# QUANTITATIVE MULTI-CONTRAST $\mu$ CT



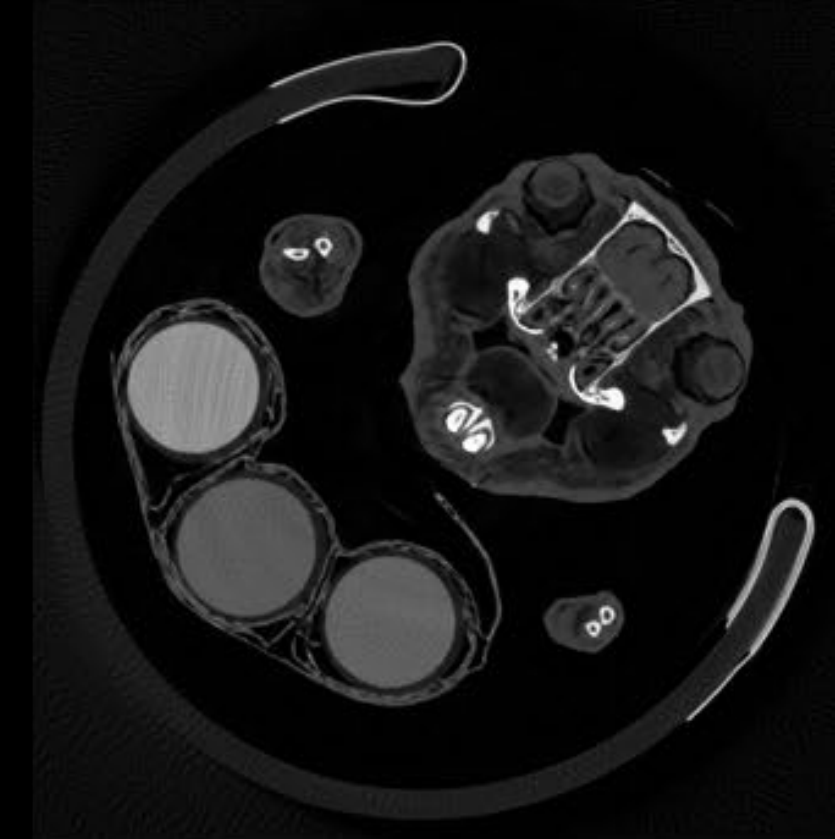
bin1 - [21, 26] keV



bin2 - [26, 33] keV



bin3 - [33, 37] keV



bin4 - [37, 42] keV



bin5 - [42, 47] keV



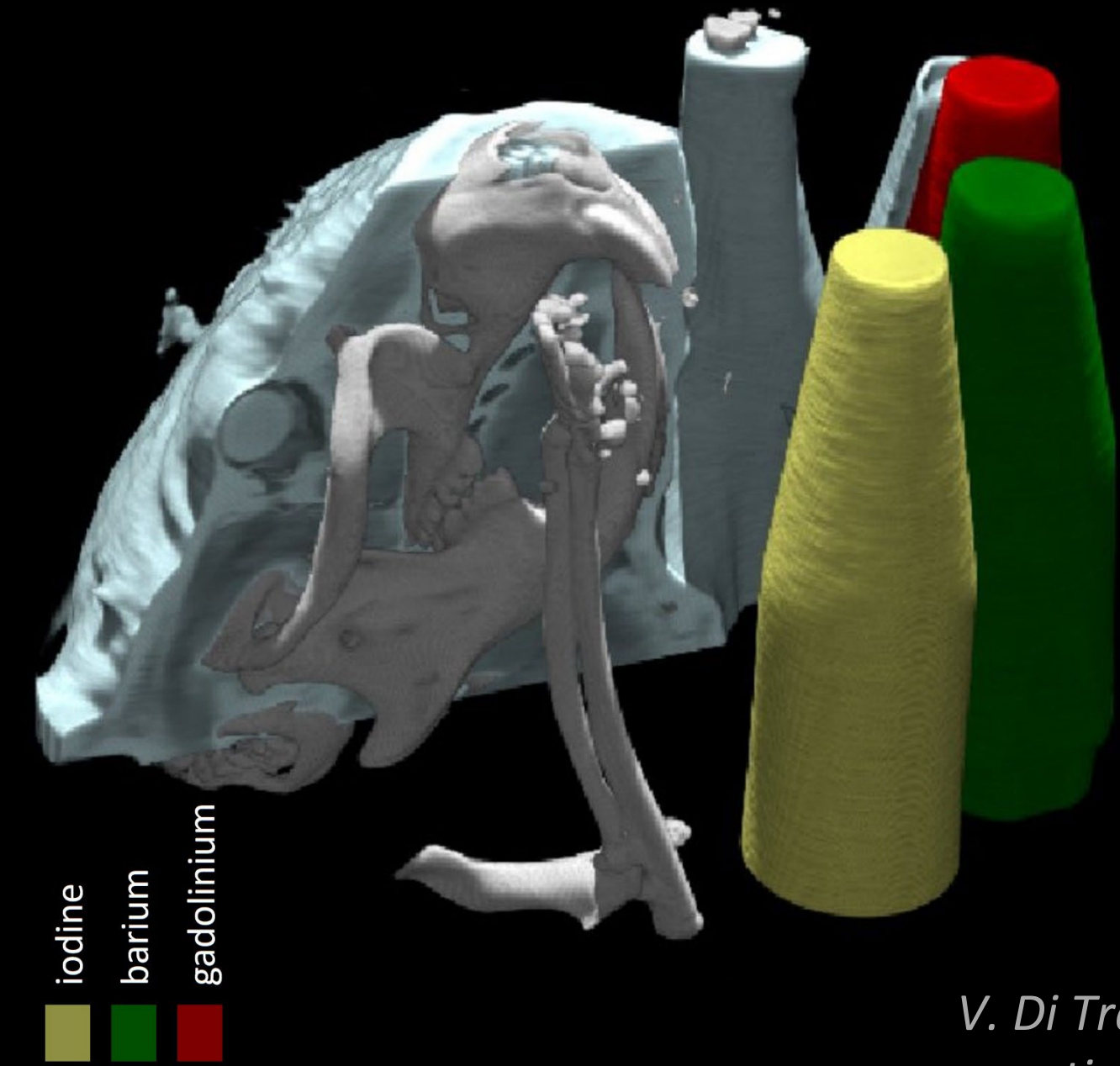
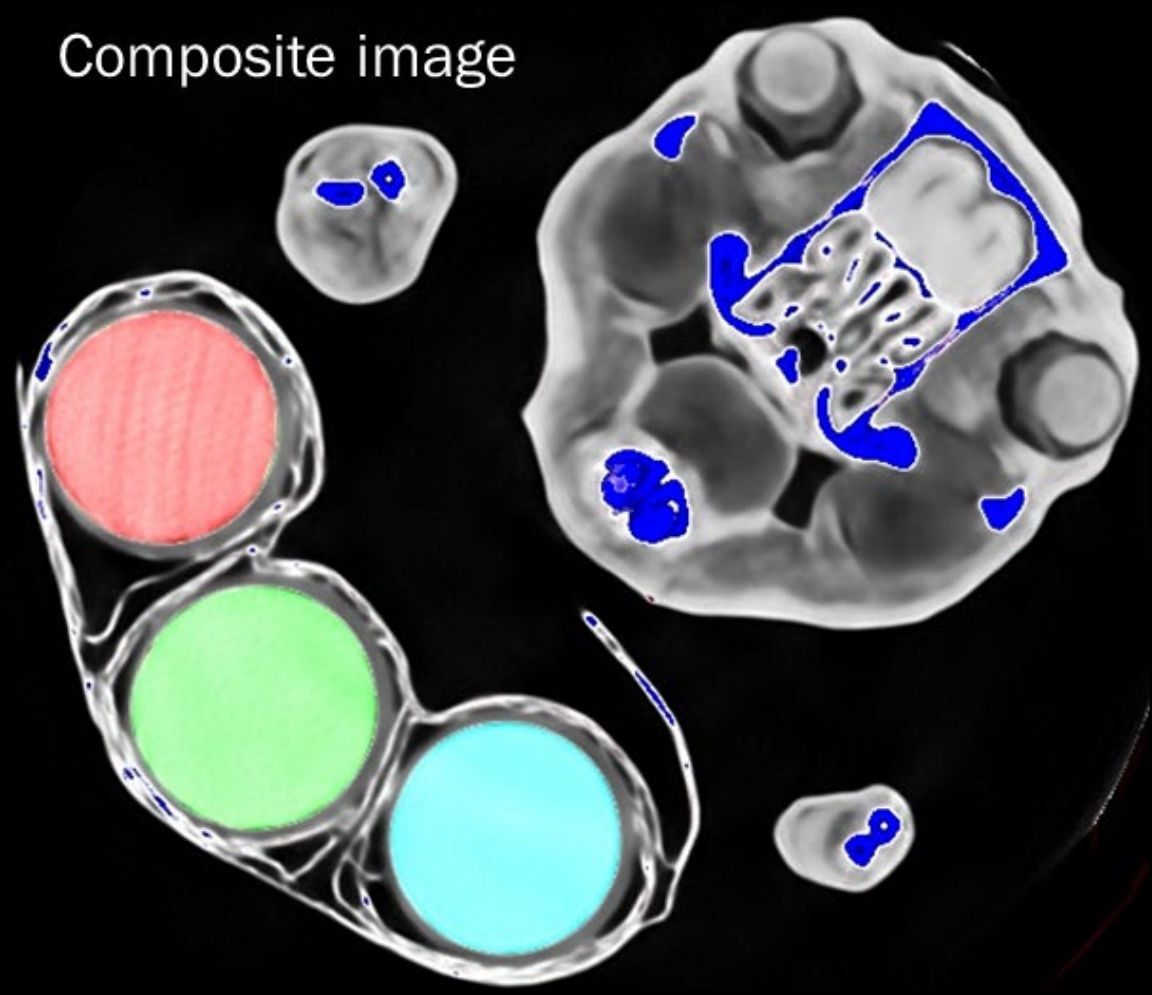
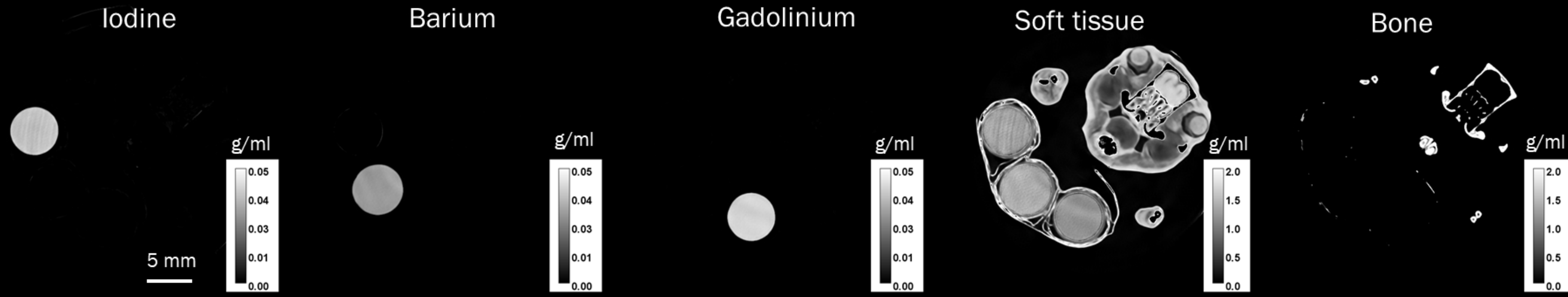
bin6 - [47, 50] keV



bin7 - [50, 57] keV

bin8 - [57,  $\infty$ ] keV

# QUANTITATIVE MULTI-CONTRAST $\mu$ CT



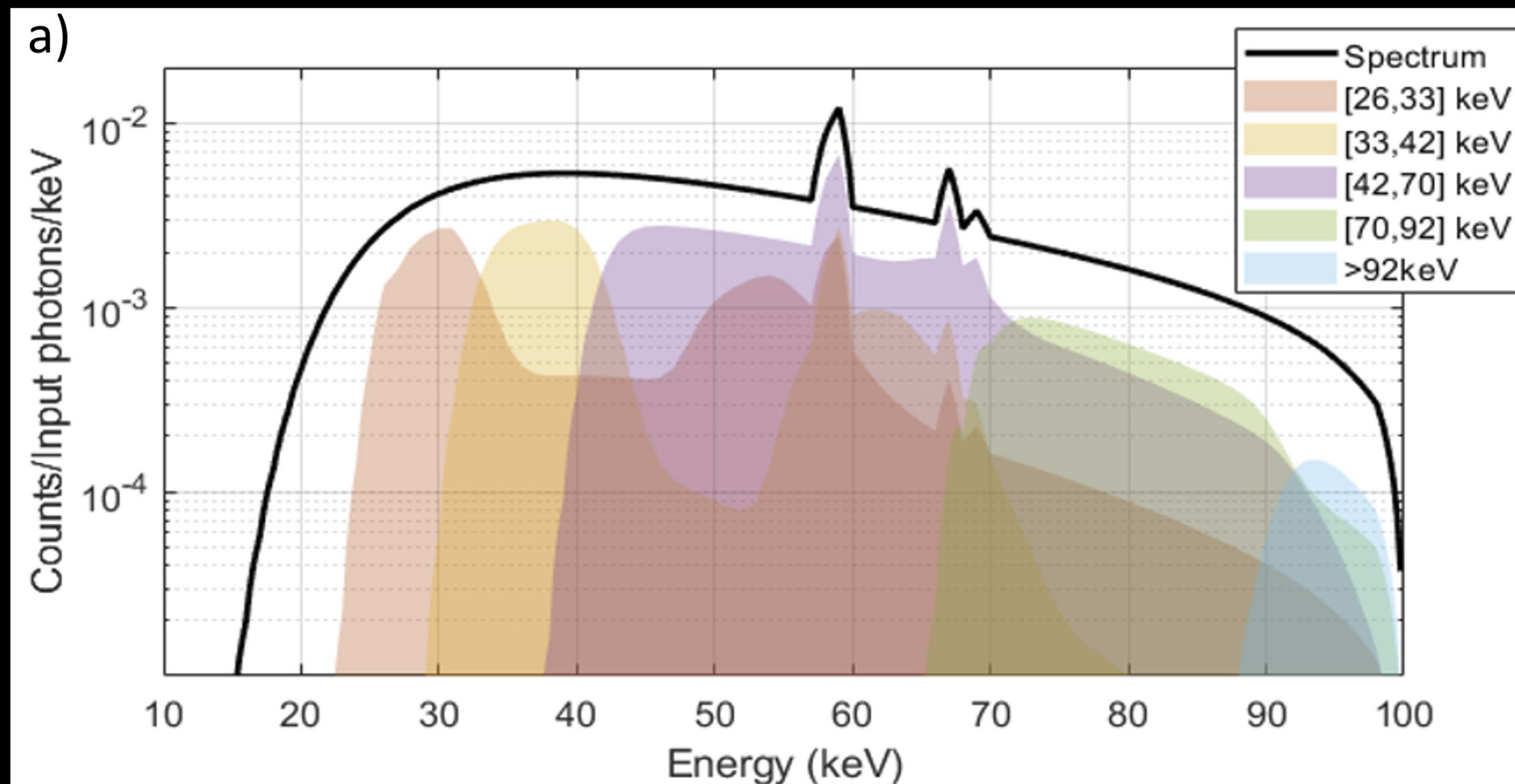
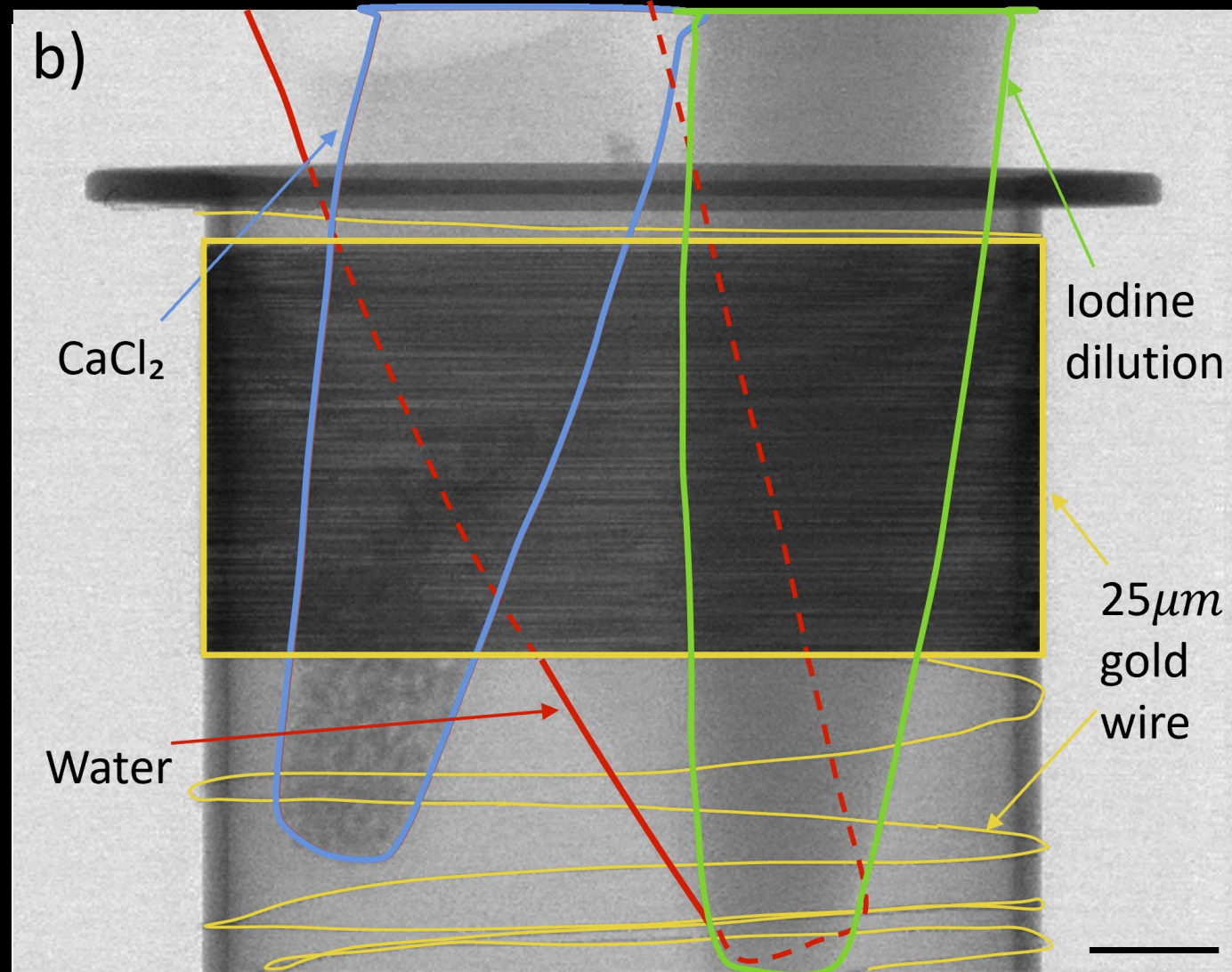
- iodine
- barium
- gadolinium
- bone
- soft tissue

	Nominal [mg/ml]	Measured [mg/ml]
Iodine	40	$37.6 \pm 0.8$
Barium	35	$30.2 \pm 0.5$
Gadolinium	39	$41.2 \pm 0.4$

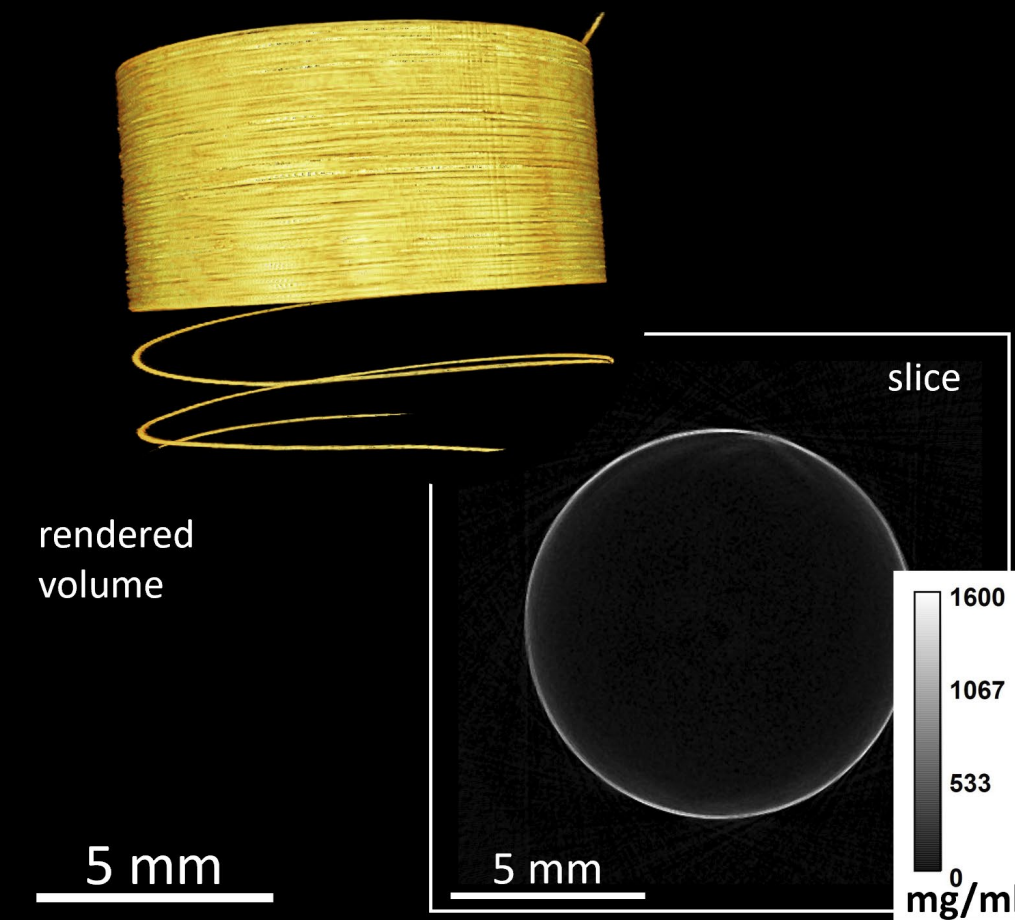
V. Di Trapani, L. Brombal, and F. Brun. "Multi-material spectral photon-counting micro-CT with minimum residual decomposition and self-supervised deep denoising." *Optics Express* 30.24 (2022): 42995-43011.



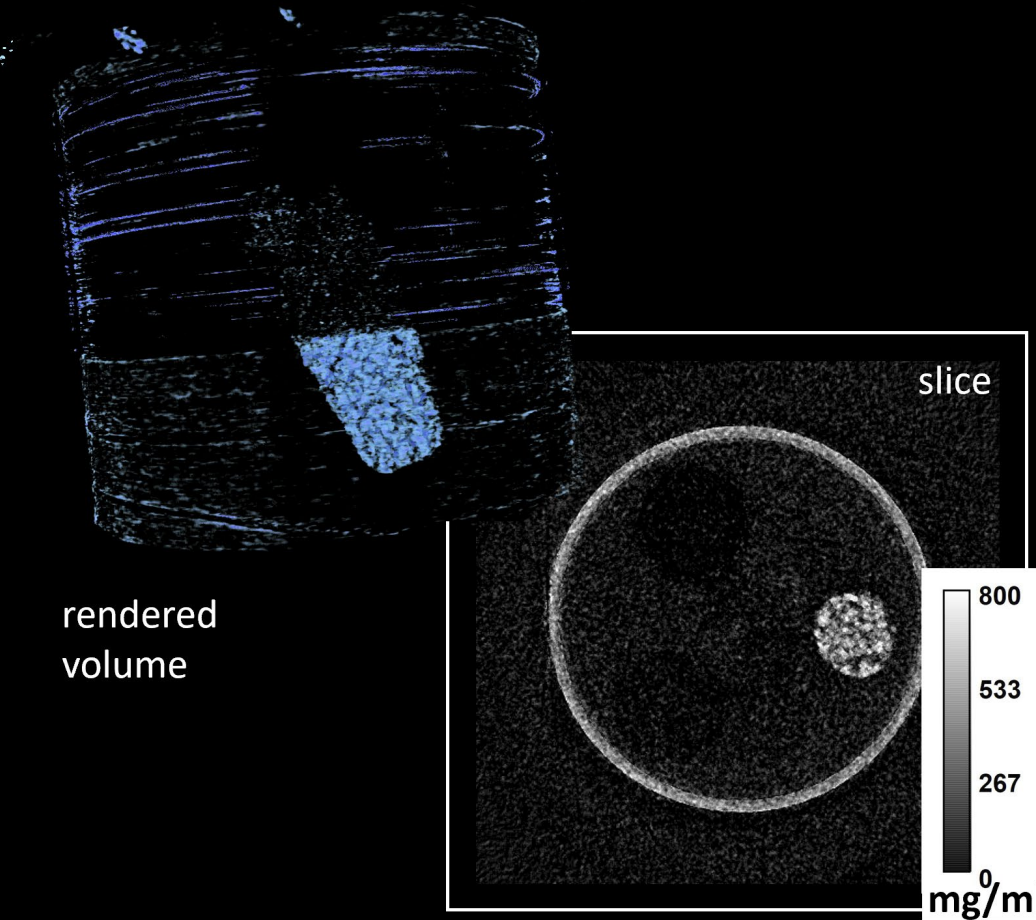
# APPLICATIONS: HIGH-ENERGY SPECTRAL IMAGING



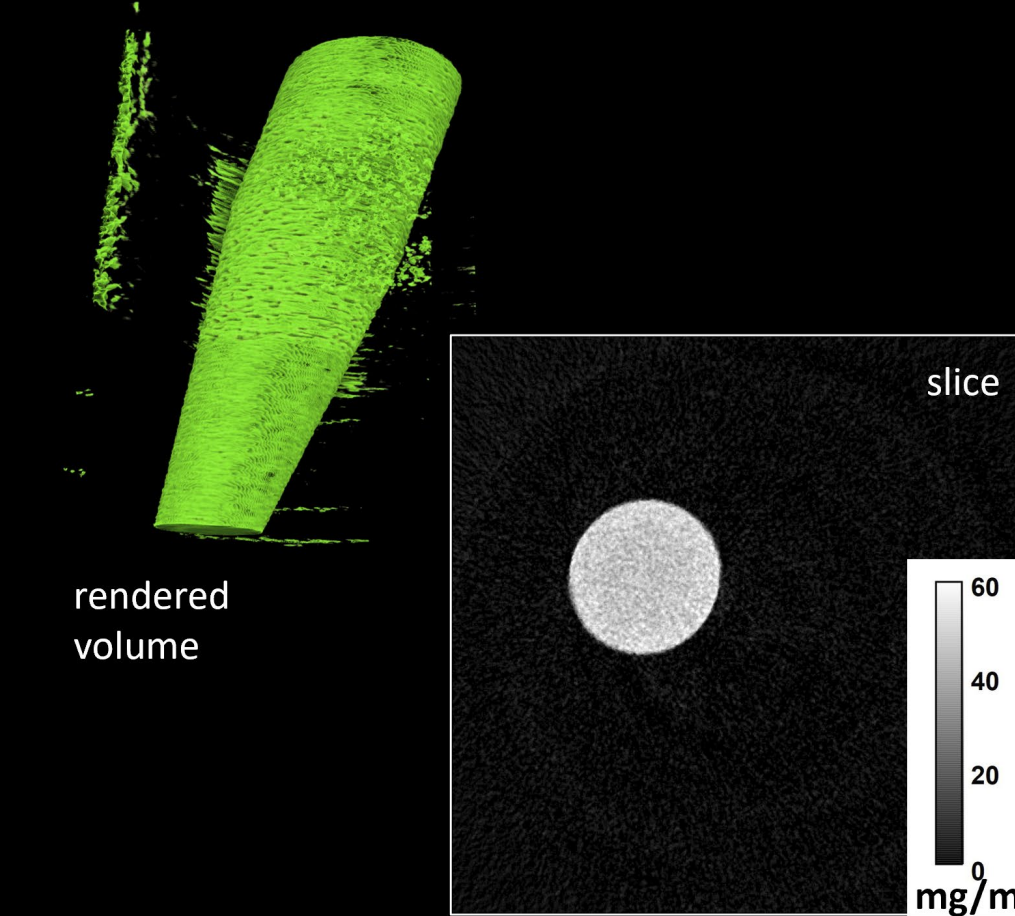
a) Gold distribution map



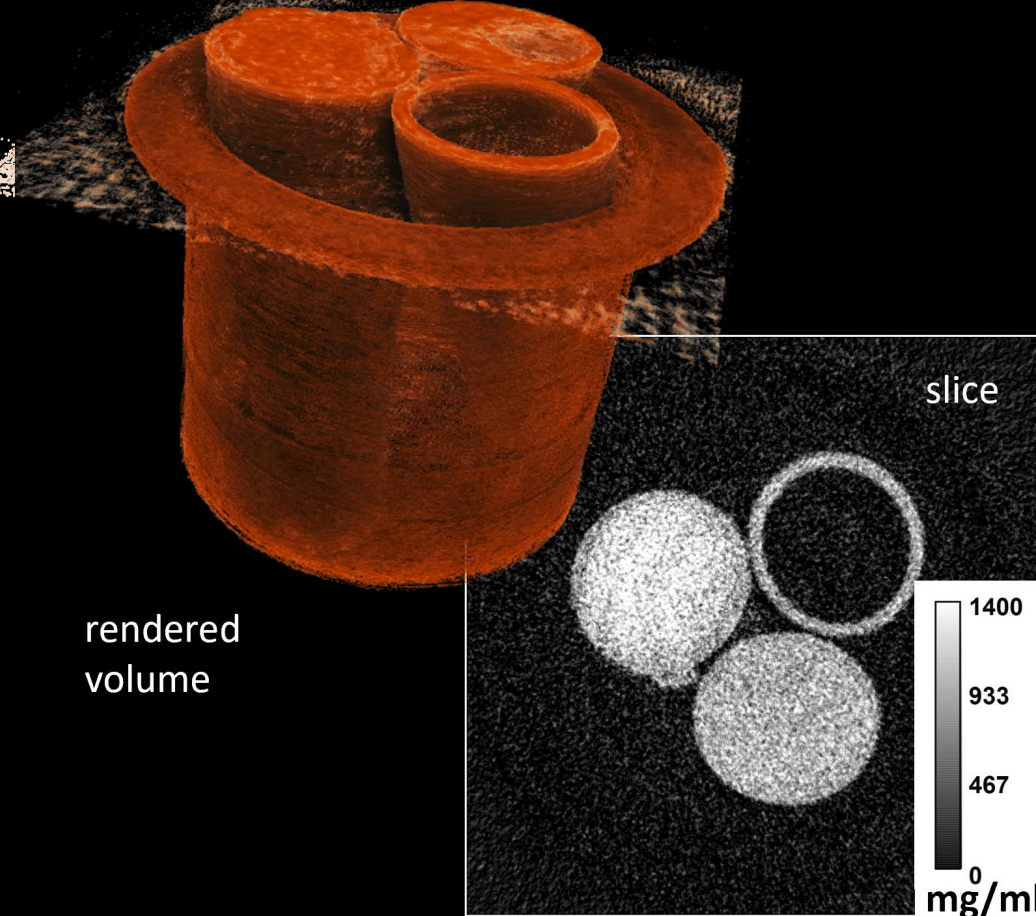
b) Calcium distribution map



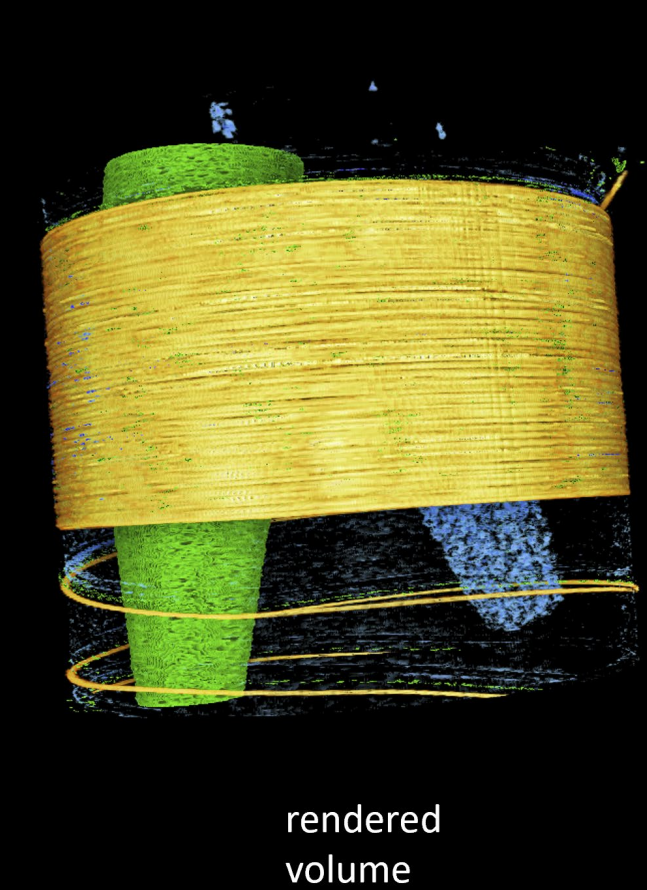
c) Iodine distribution map

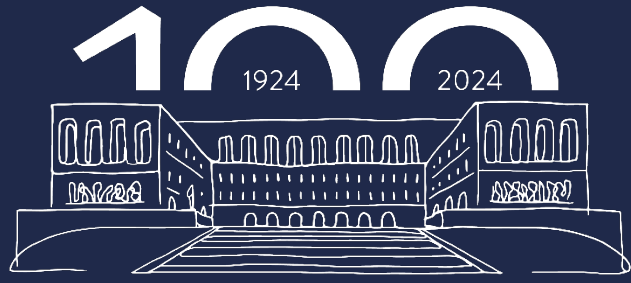


d) Water distribution map



e) Gold + Calcium + Iodine





UNIVERSITÀ  
DEGLI STUDI  
DI TRIESTE

# OUTLINE

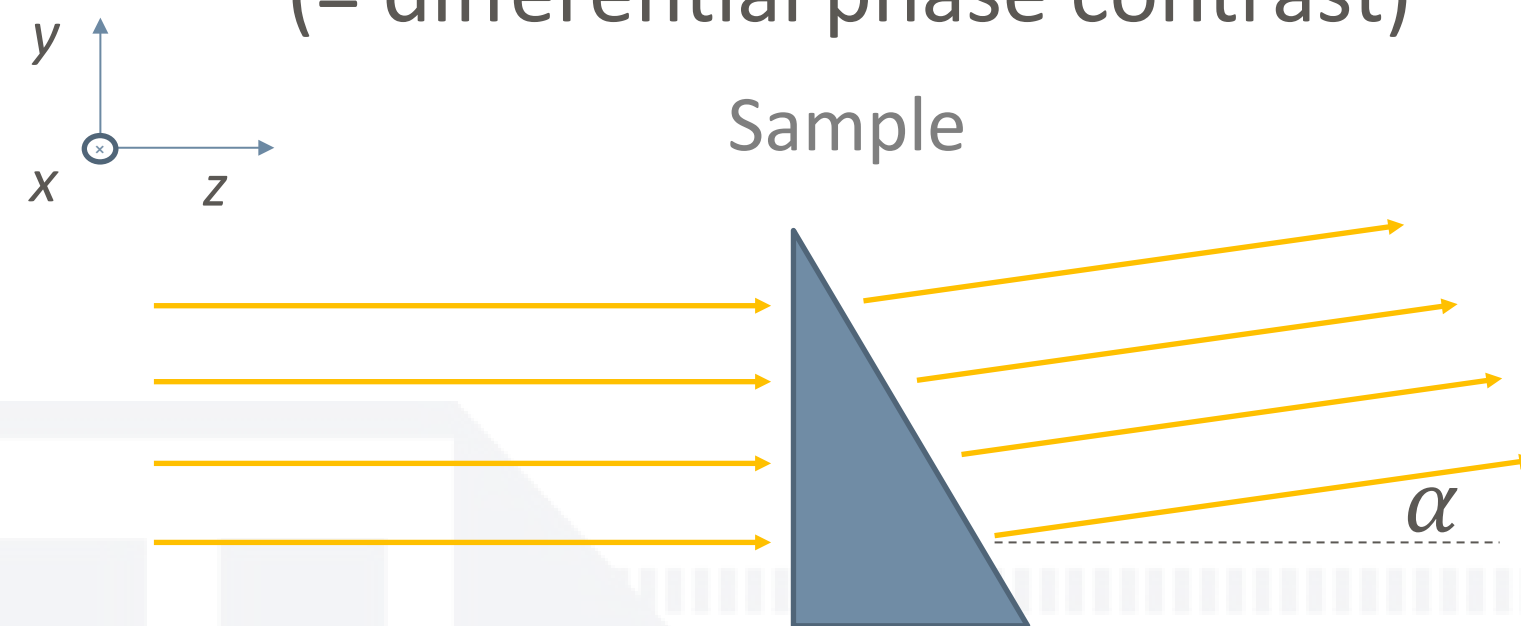
- X-ray imaging fundamentals
- Spectral imaging
- **Phase-contrast imaging**
- Spectral phase-contrast imaging

# PHASE EFFECTS

A naïve interpretation

## Refraction

(= differential phase contrast)

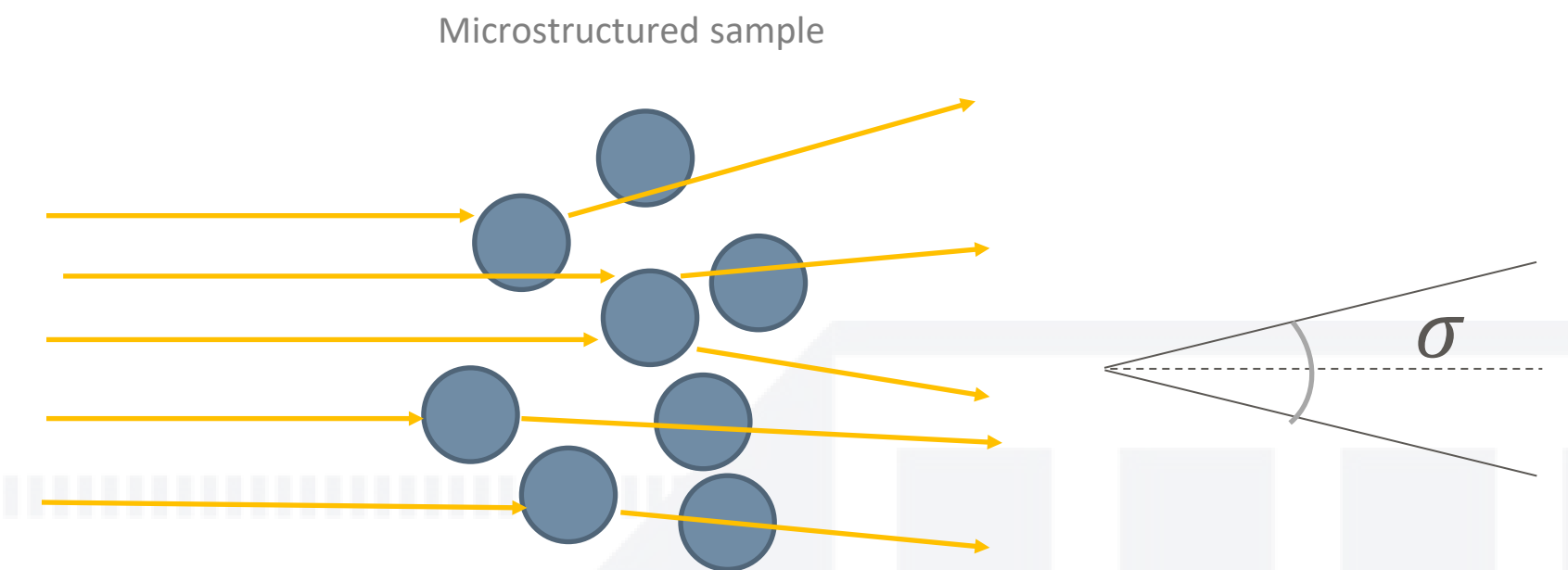


$$\alpha = -\nabla_{xy} \int \delta(x, y, z; E) dz$$

- Within the ray-optical approximation phase effects = *refraction*
- Refraction is proportional to the gradient of  $\delta \rightarrow$  strong at the edges
- Refraction angles range 1-100  $\mu\text{rad}$

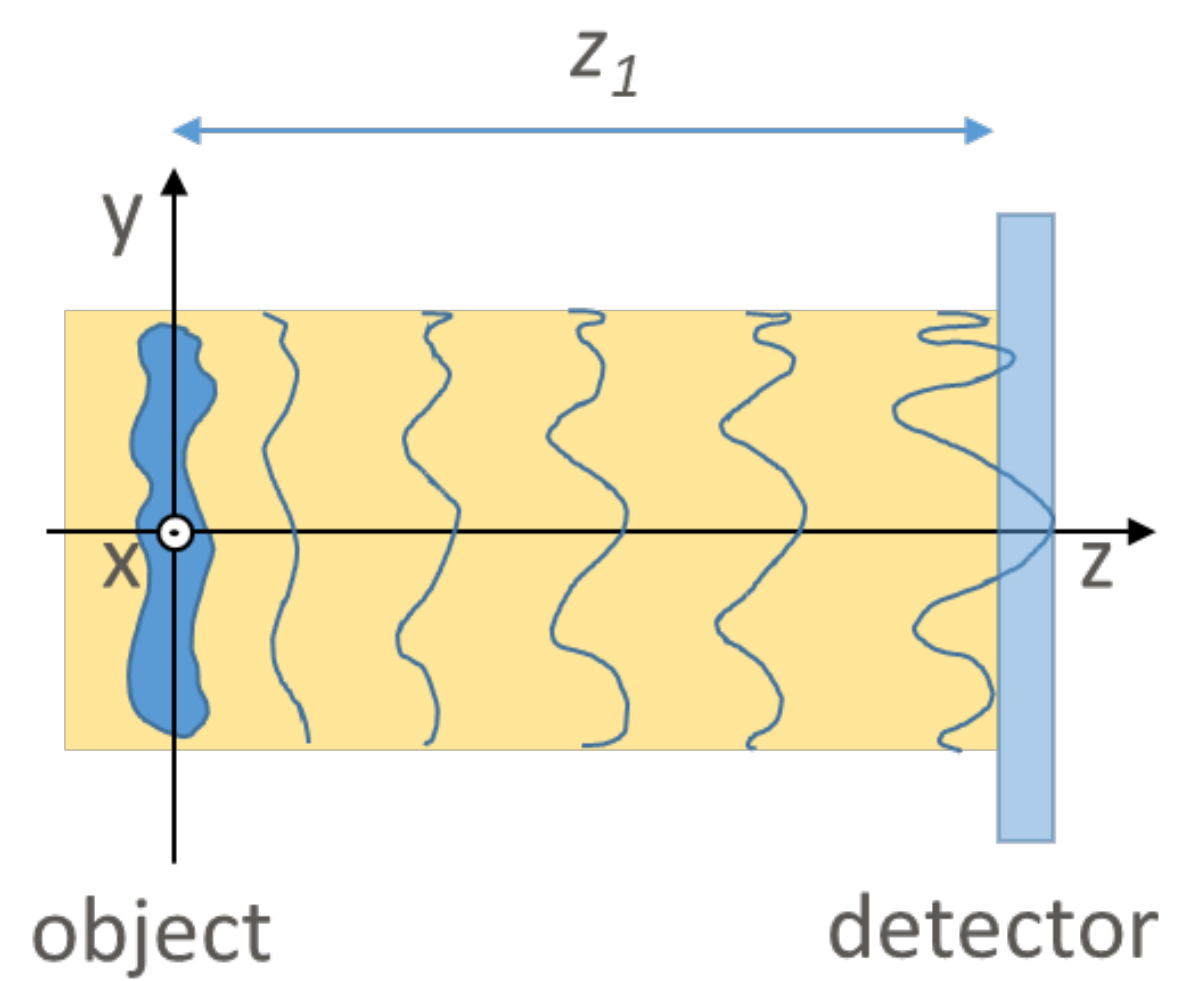
## Ultra-small angle scattering

(= dark field)

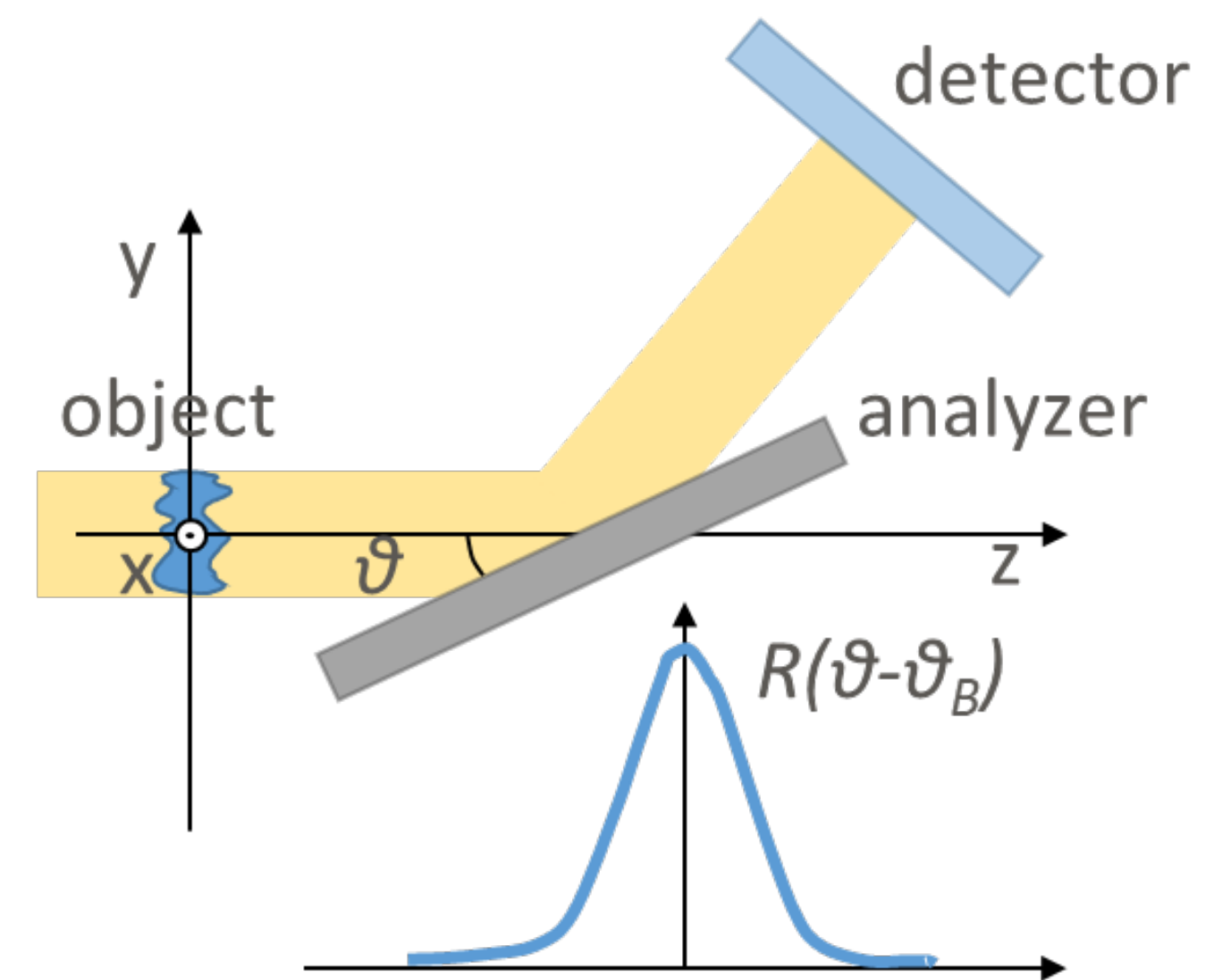


- In microstructured samples *multiple-refraction* occurs, causing a diffusion of the beam in the range 1-100  $\mu\text{rad}$
- The “amount of diffusion”, i.e. scattering signal, depends on sample’s properties at a scale smaller than the system’s spatial resolution

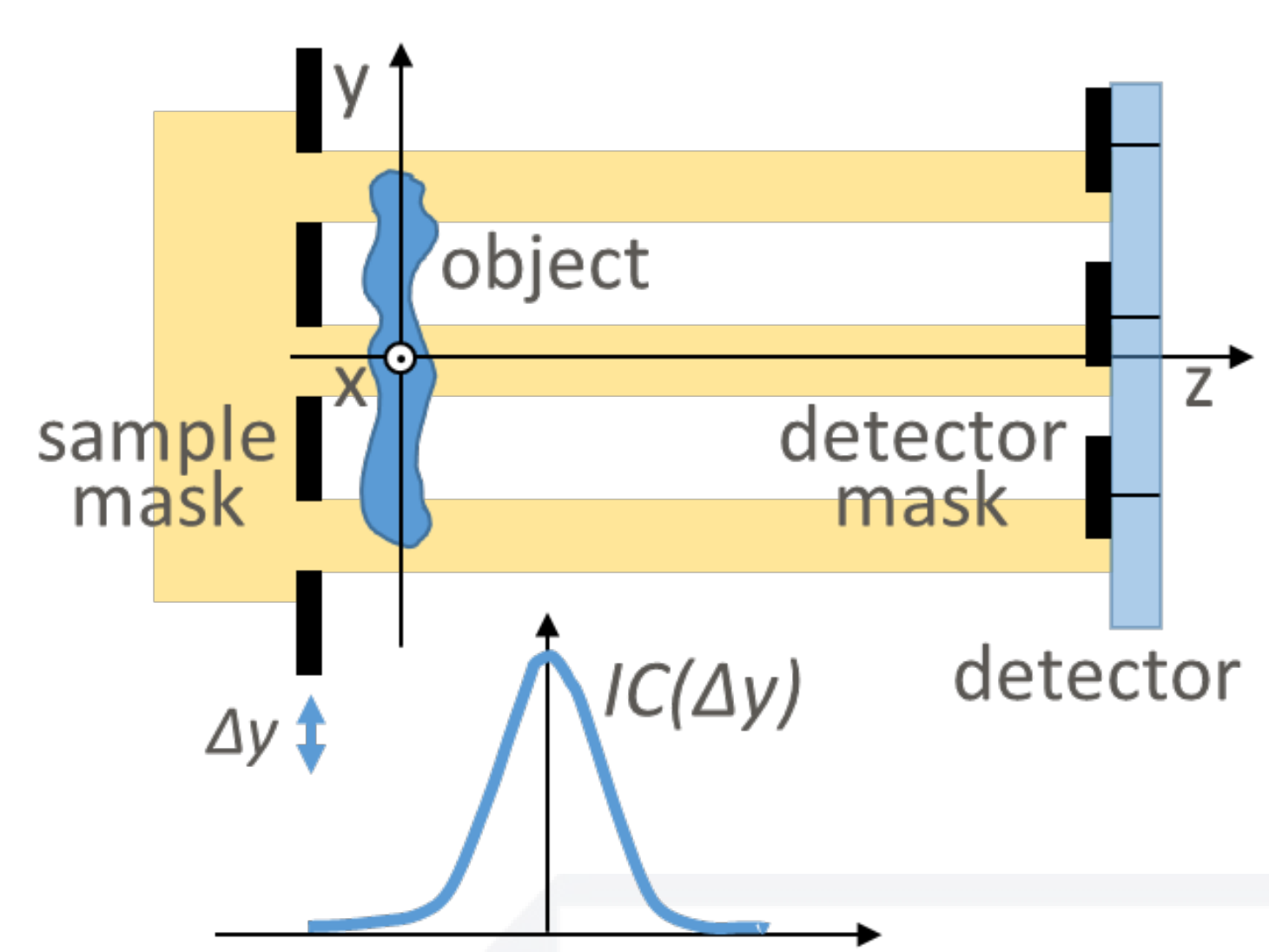
### PROPAGATION-BASED



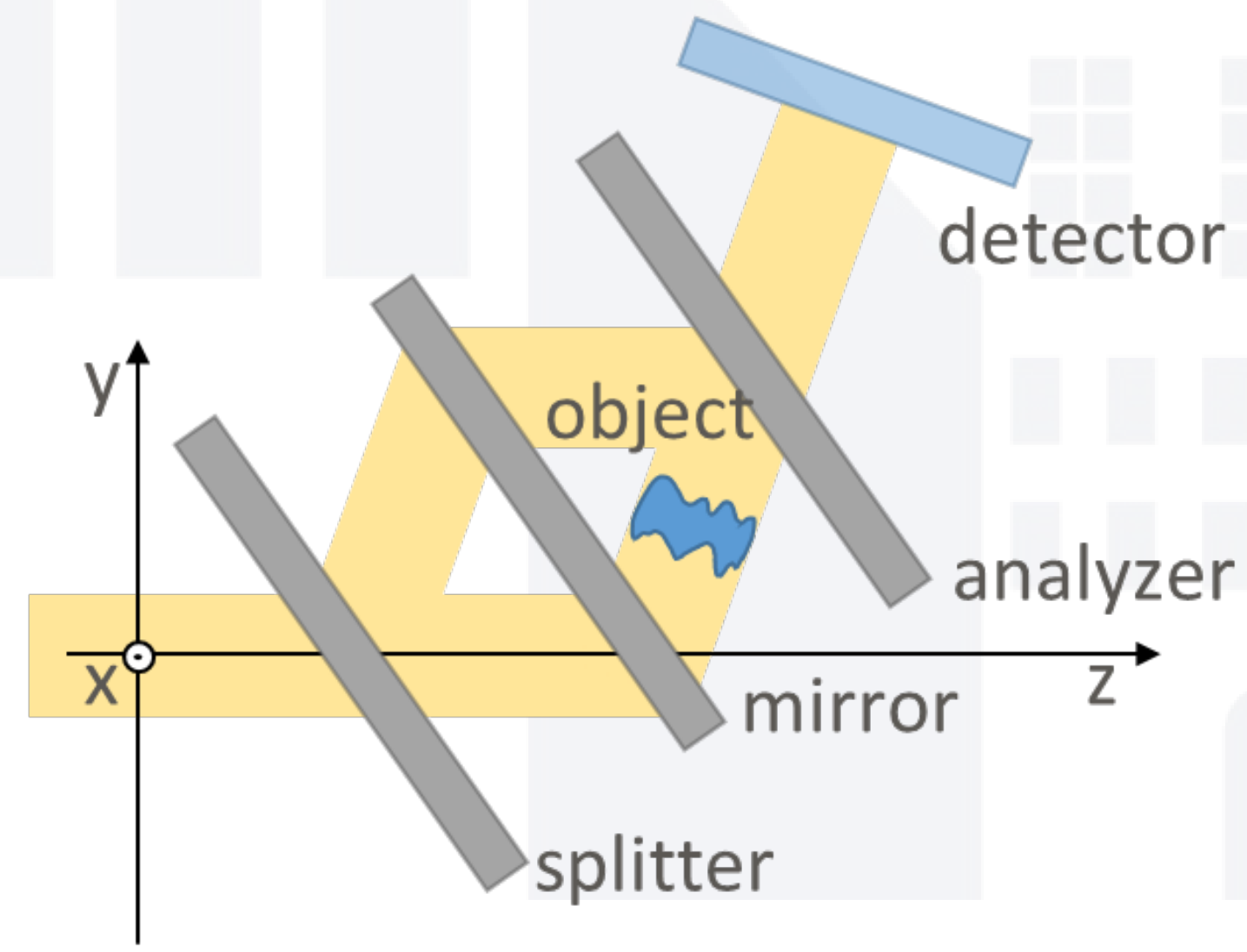
### ANALYZER-BASED



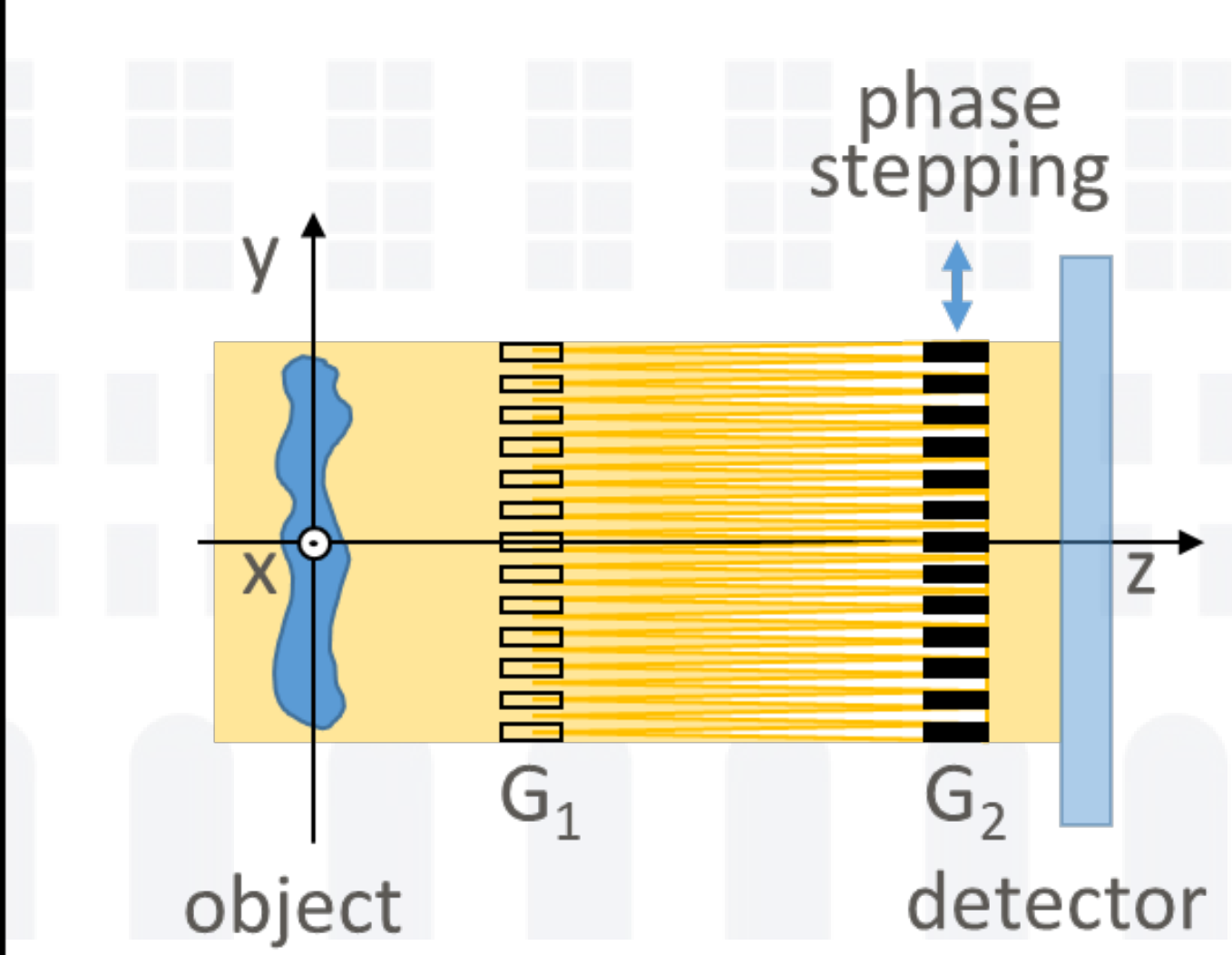
### EDGE ILLUMINATION



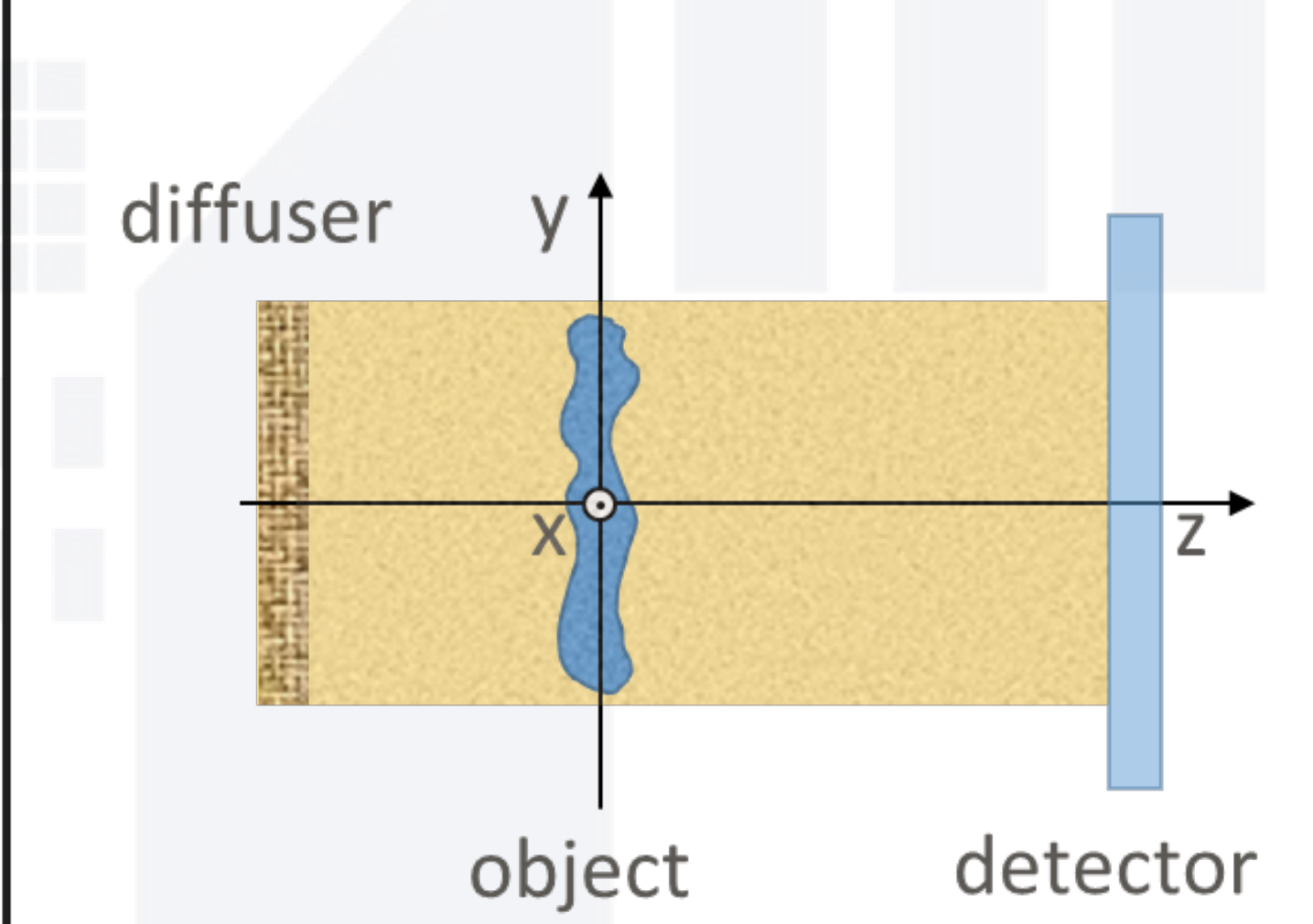
### INTERFEROMETRY



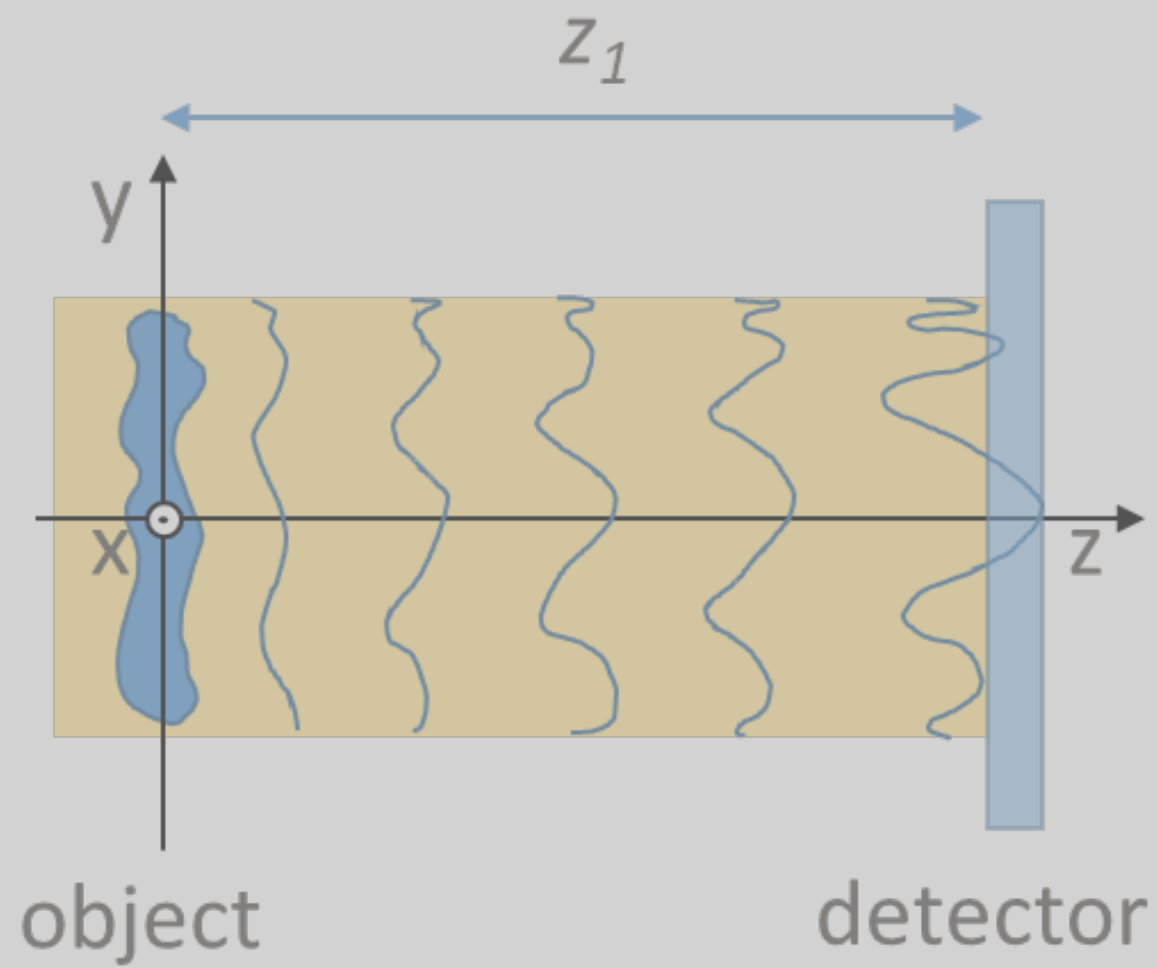
### GRATING INTERFEROMETRY



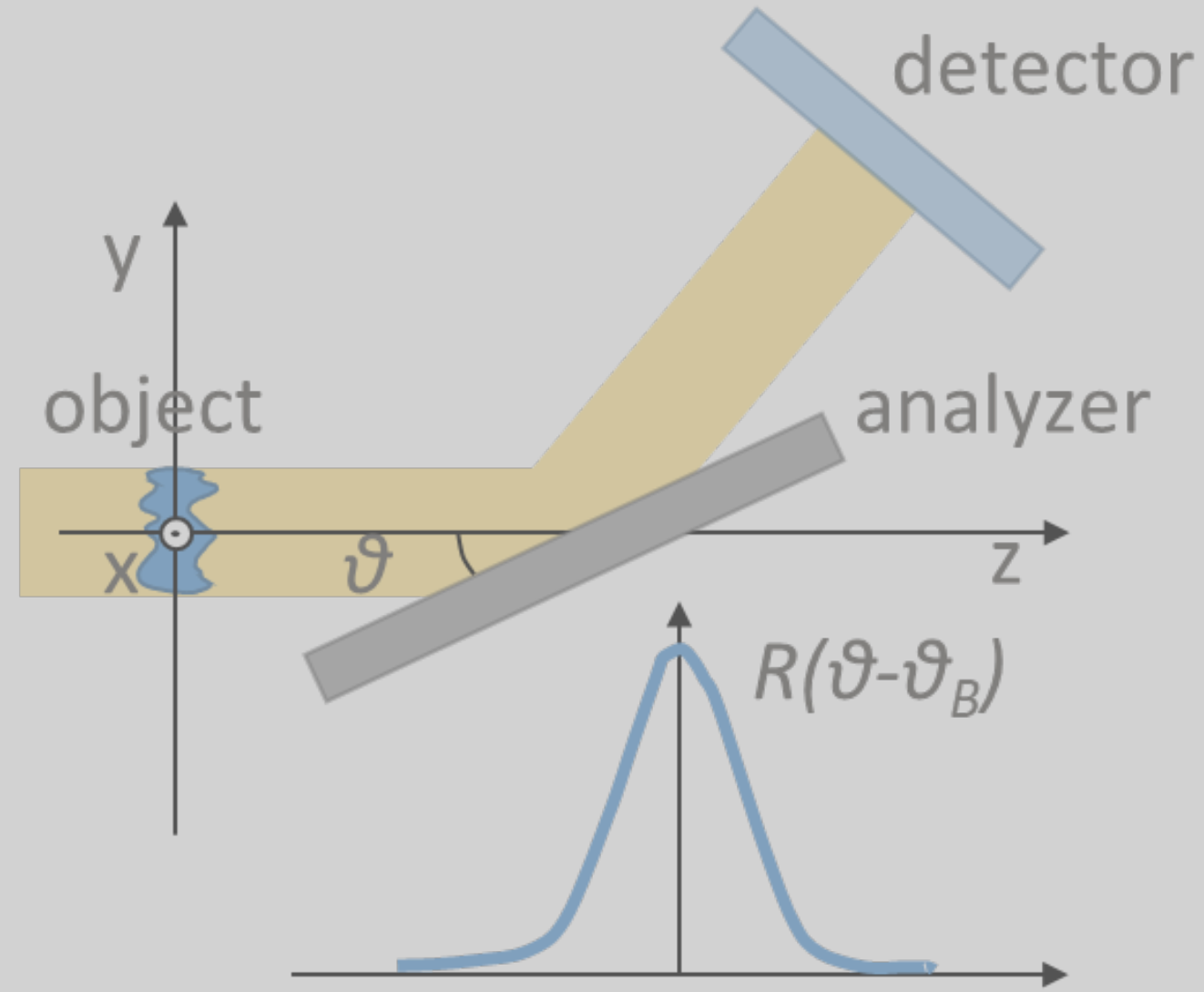
### SPECKLE IMAGING



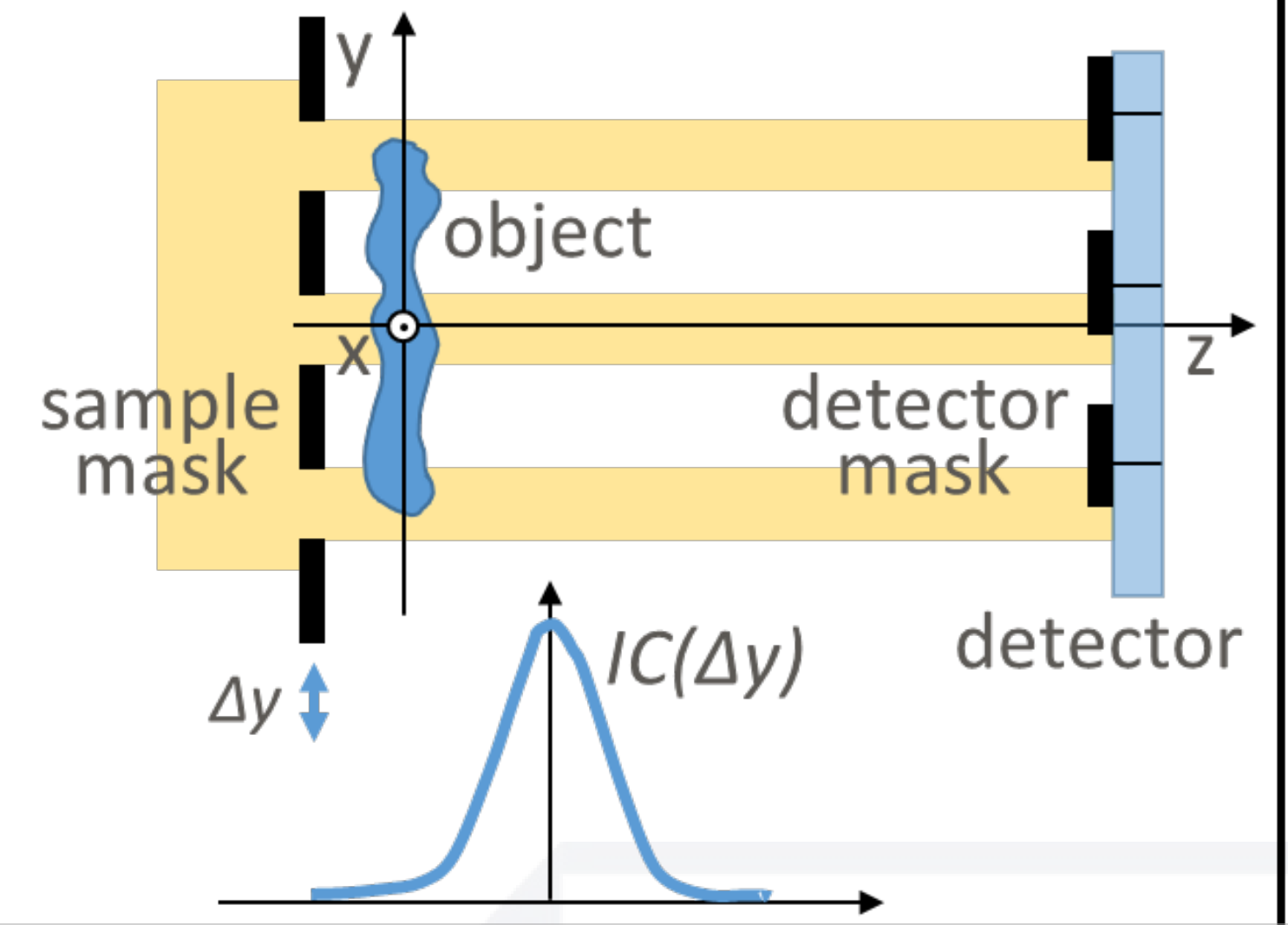
### PROPAGATION-BASED



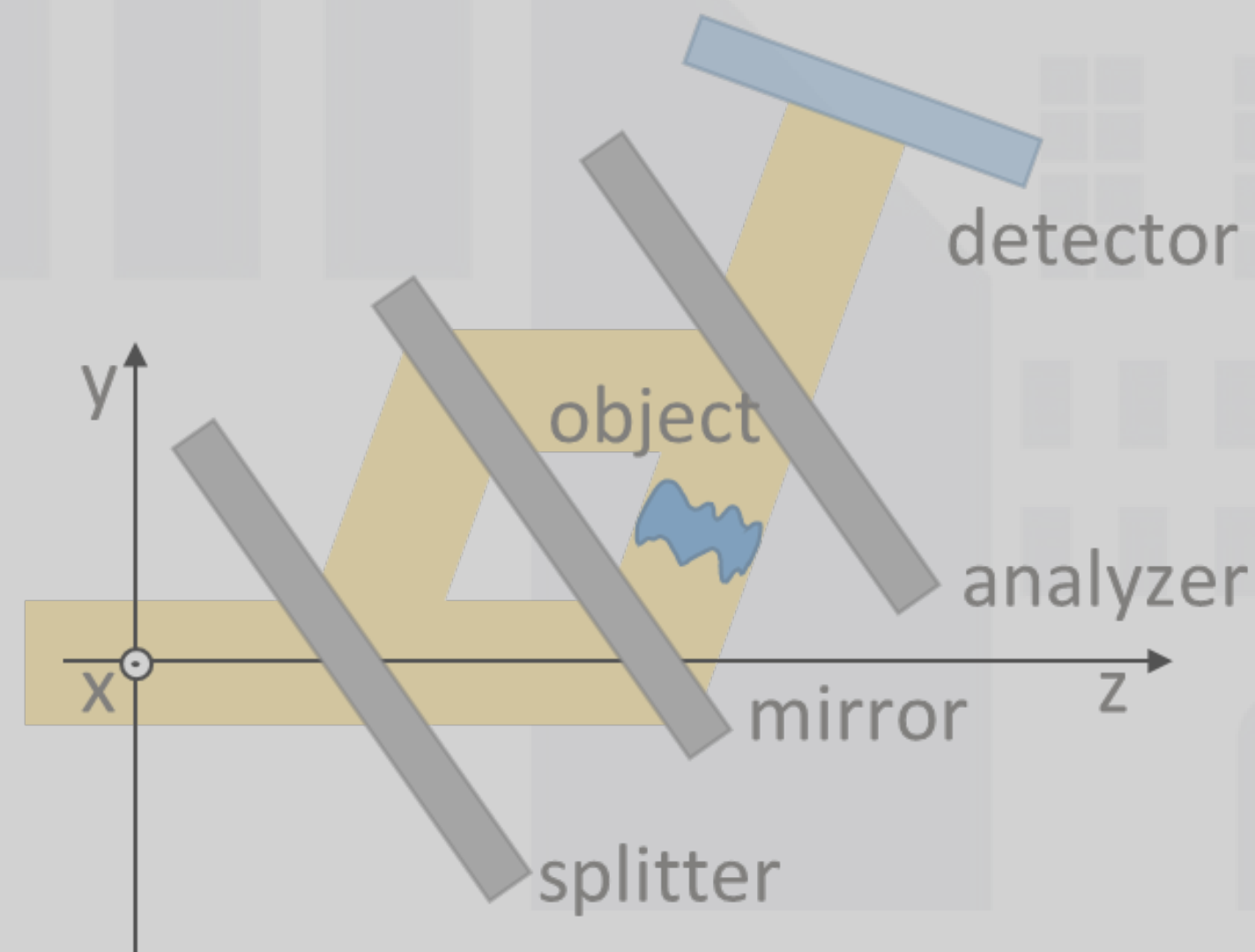
### ANALYZER-BASED



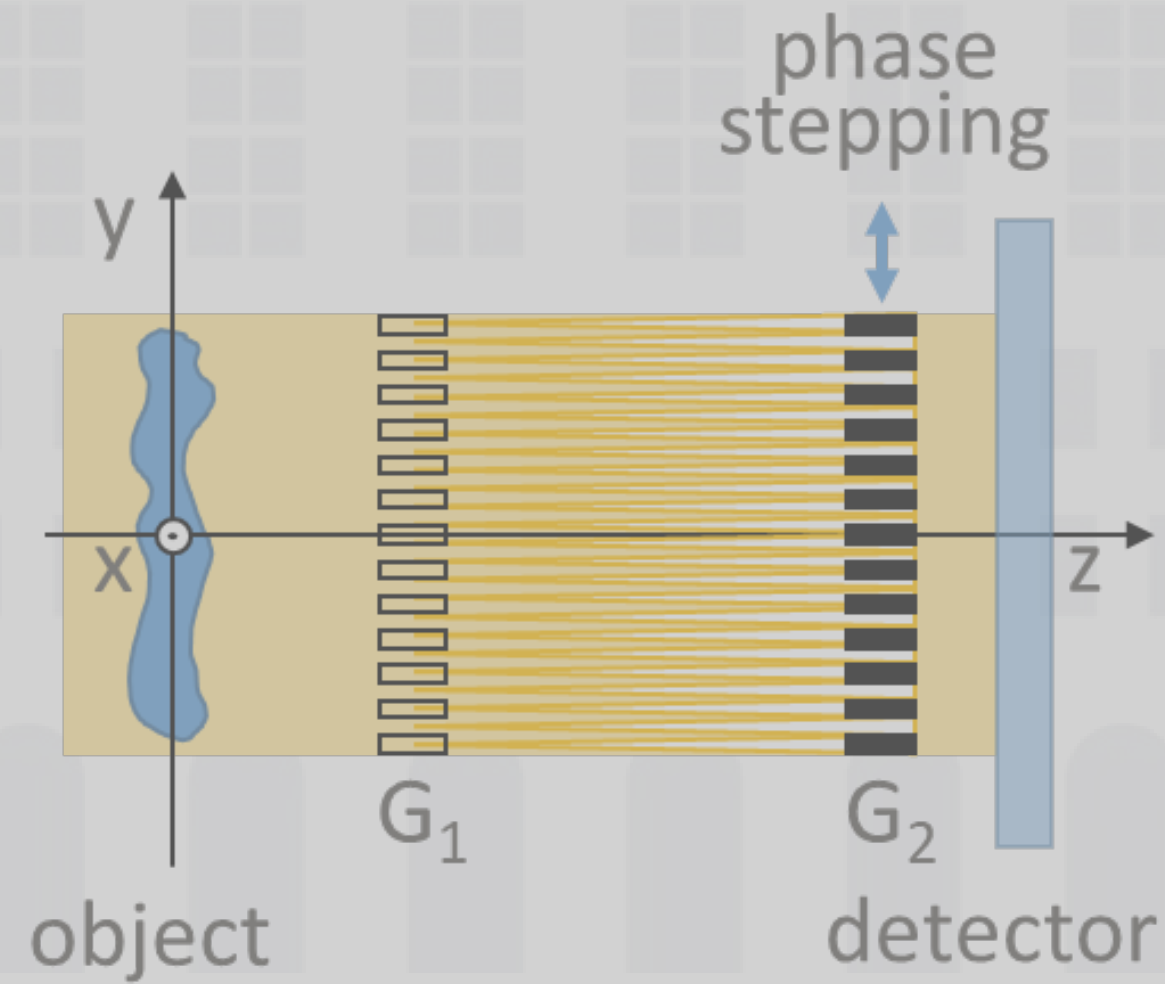
### EDGE ILLUMINATION



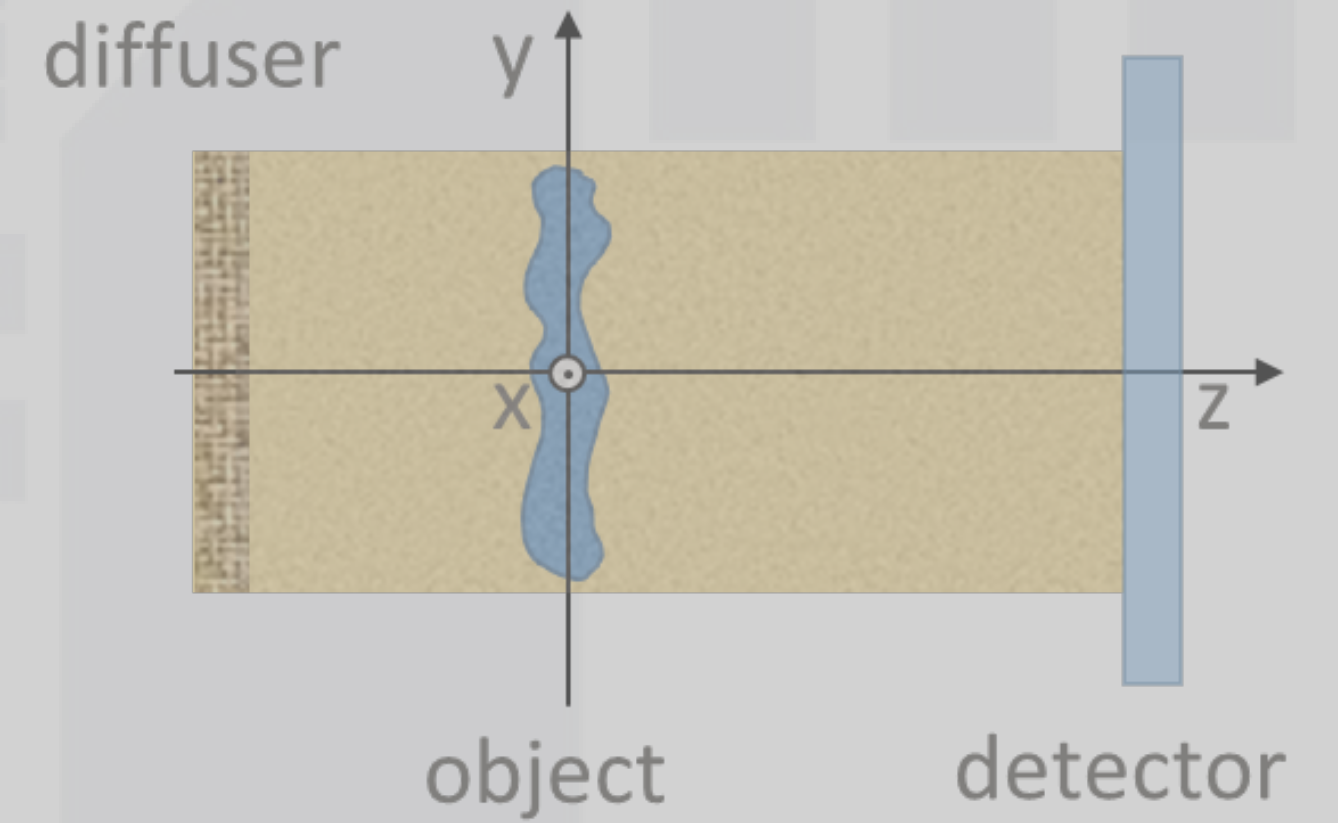
### INTERFEROMETRY



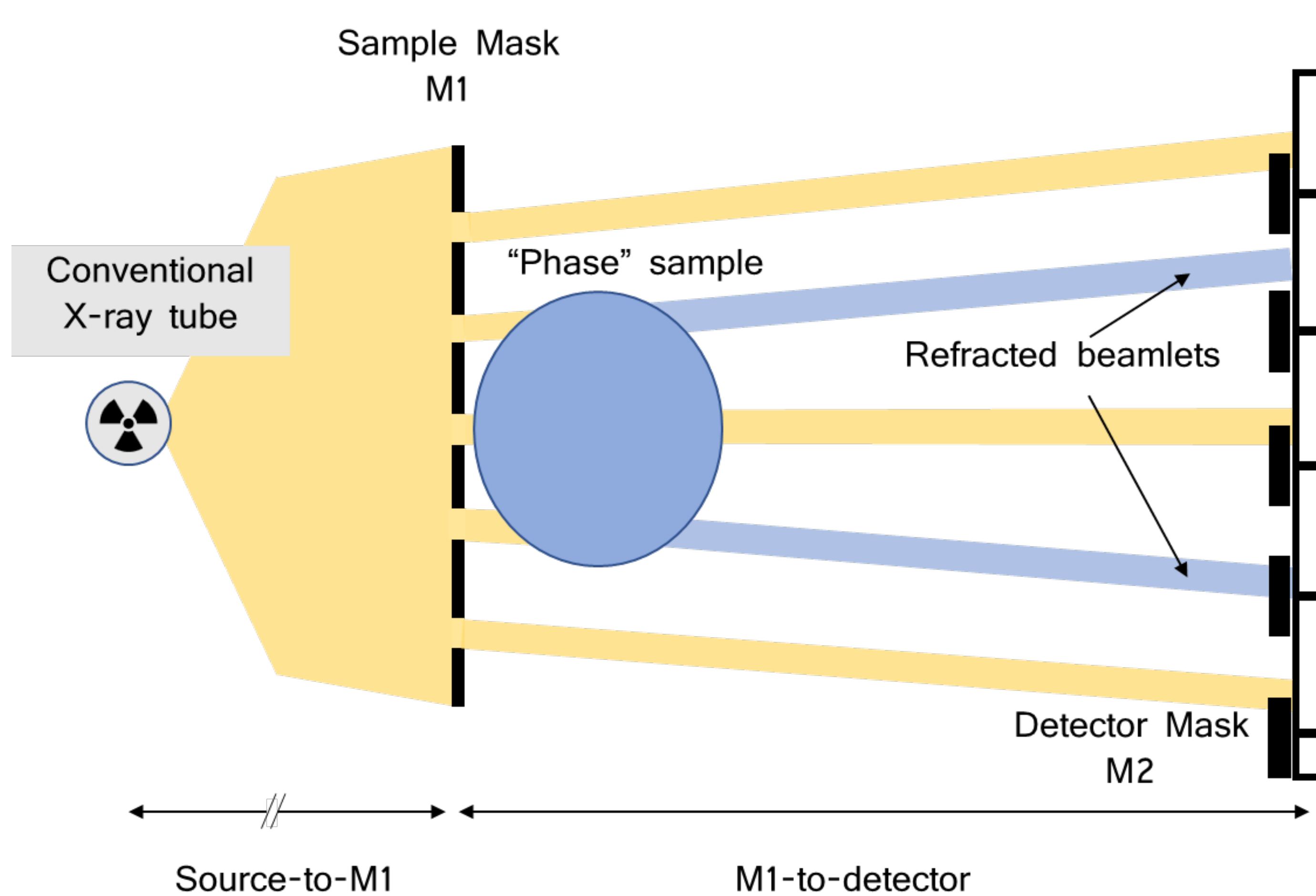
### GRATING INTERFEROMETRY



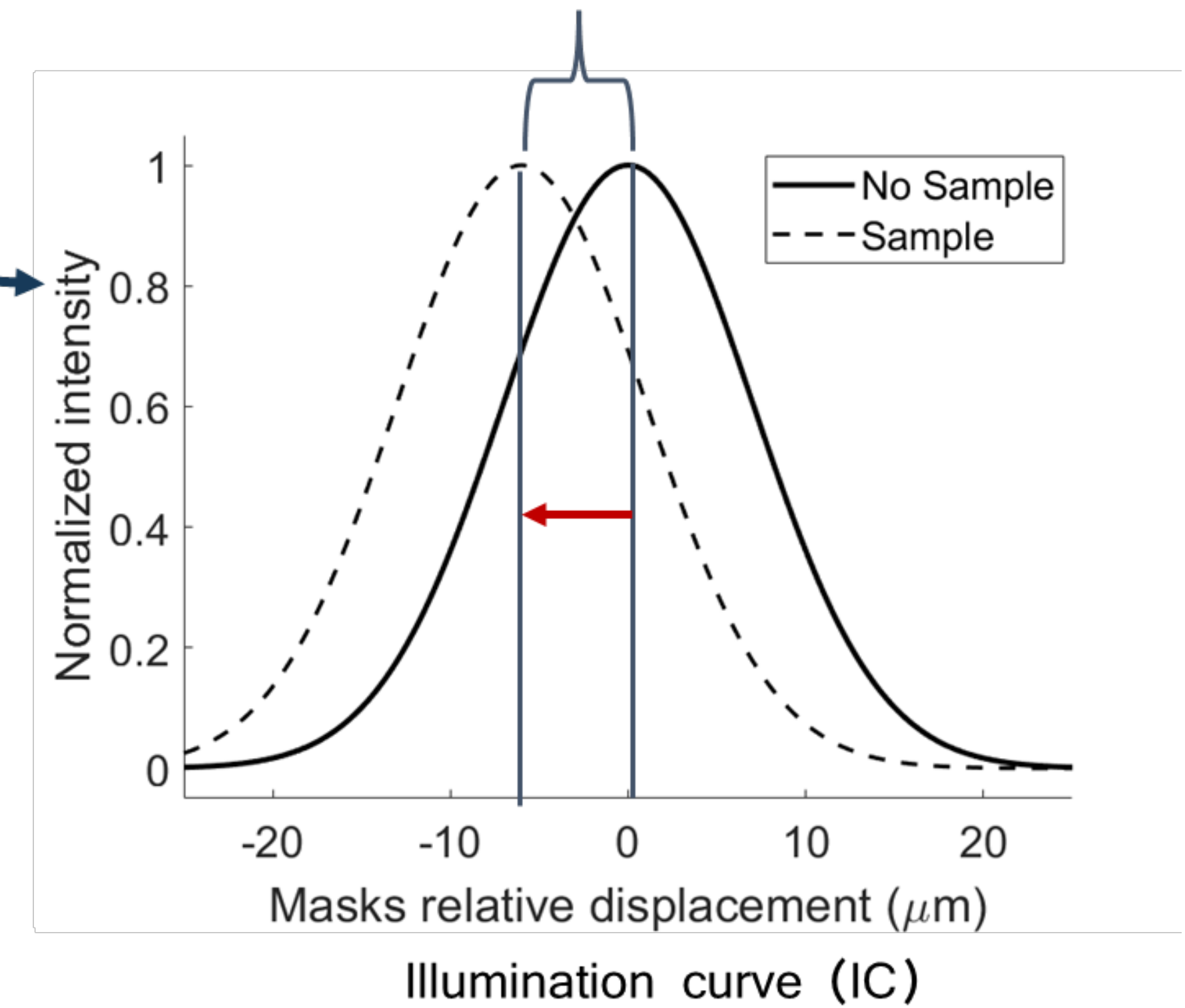
### SPECKLE IMAGING



# EDGE ILLUMINATION: HOW IT WORKS

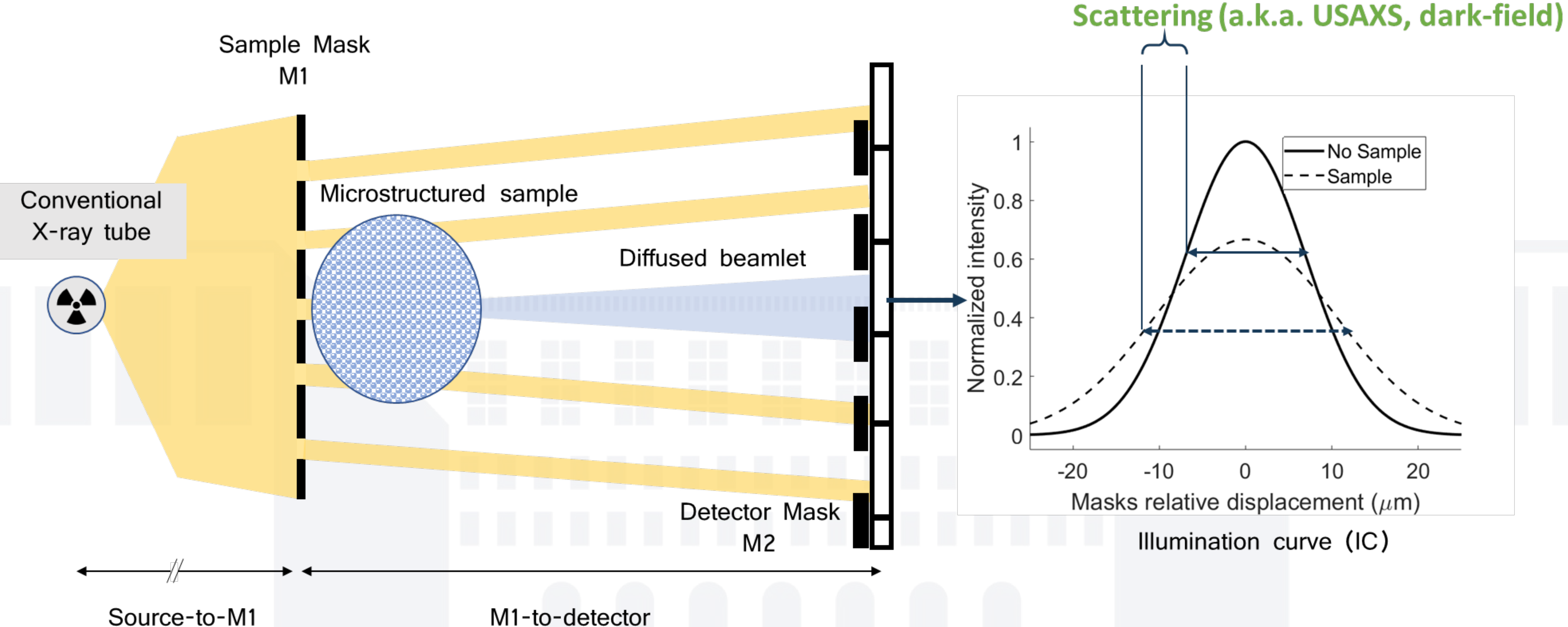


## Refraction (due to phase-shift)

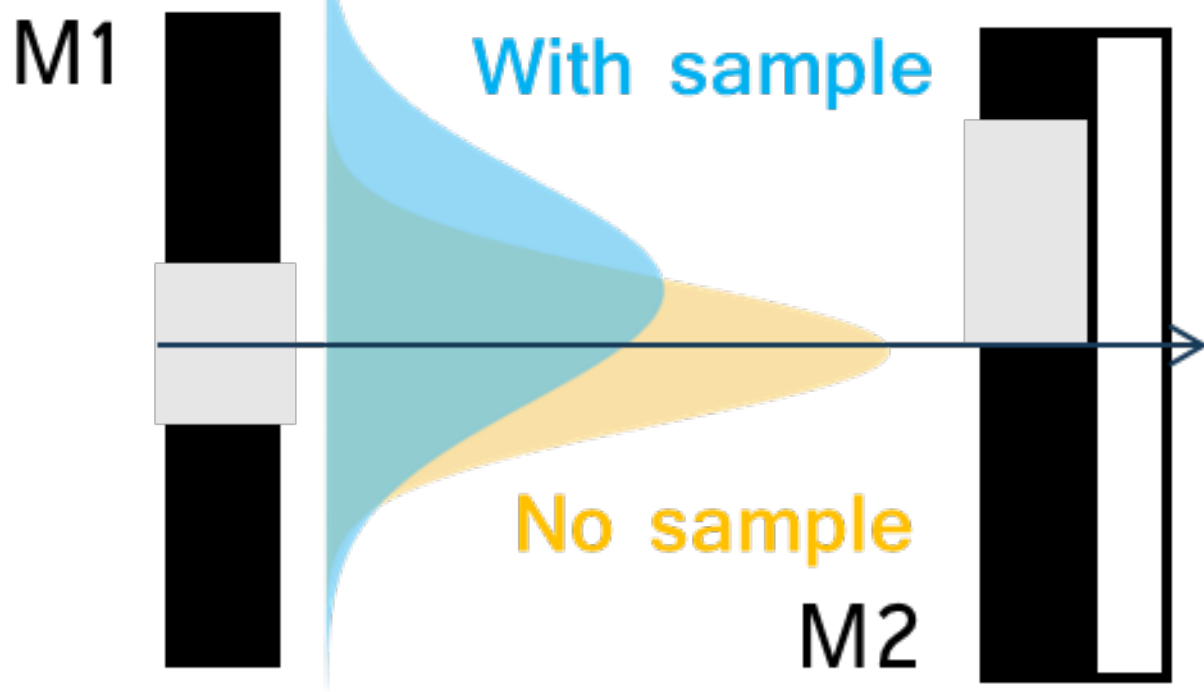


Olivo, Alessandro. "Edge-illumination x-ray phase-contrast imaging." *Journal of Physics: Condensed Matter* 33.36 (2021): 363002.

# EDGE ILLUMINATION: HOW IT WORKS



# EDGE ILLUMINATION: SIGNAL RETRIEVAL

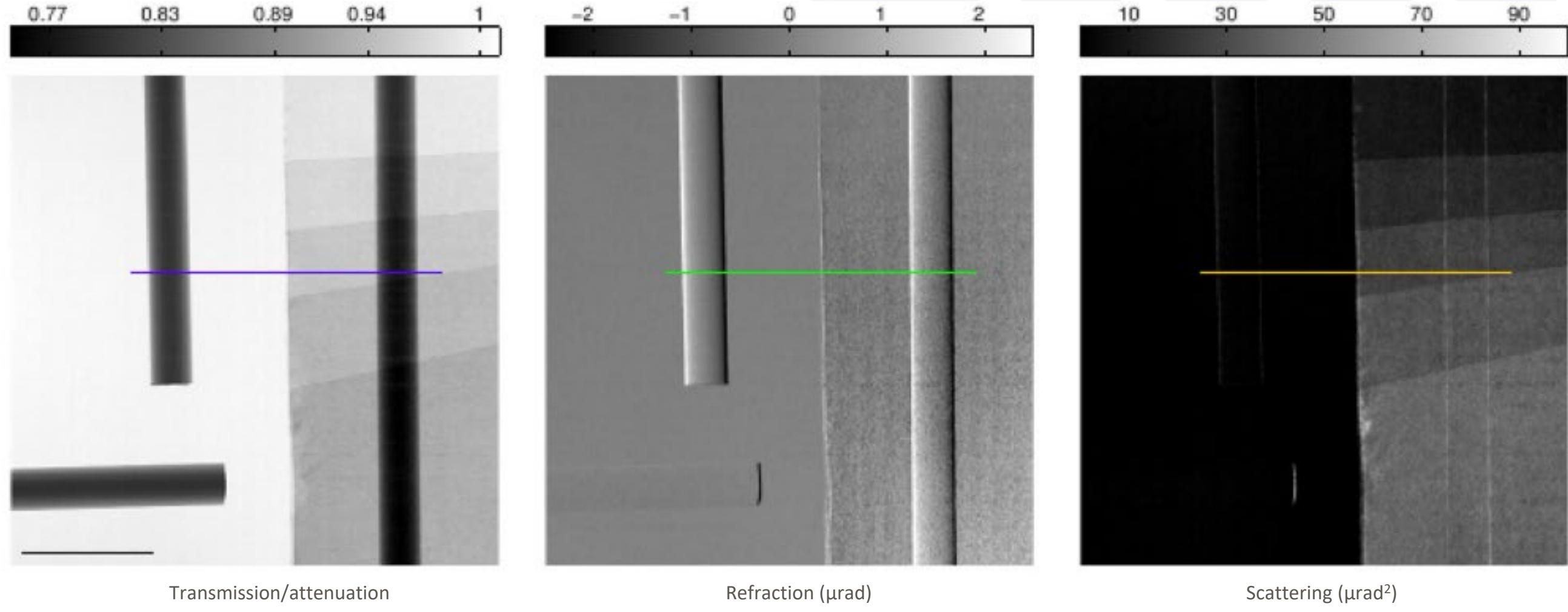
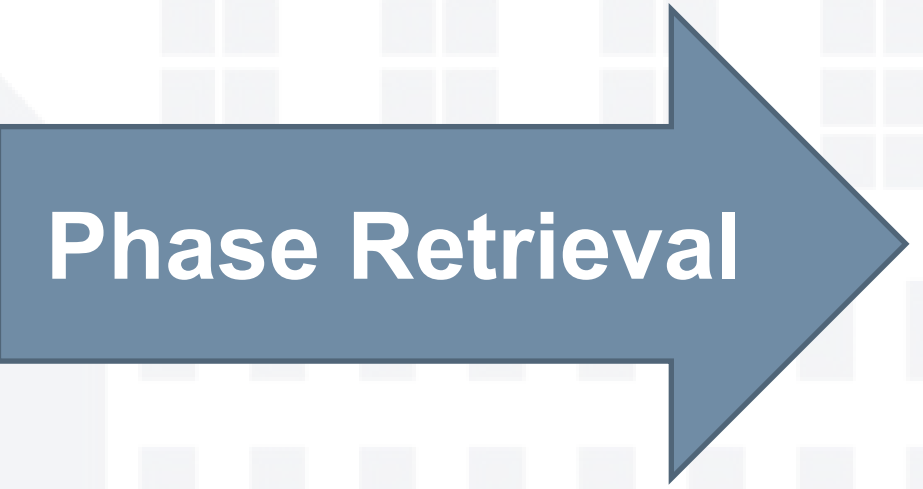


## 2/3 parametric images

- Attenuation  $\mu \propto \beta \rightarrow$  conventional X-ray imaging
- Refraction angle  $\alpha \propto \nabla \delta \rightarrow$  refraction/phase maps (via integration)
- Scattering  $\sigma \rightarrow$  Dark-field or USAXS imaging

$$I(x) = (S * IL)(x - \Delta x)t$$

2/3 (or more) input images at different Positions on the IC



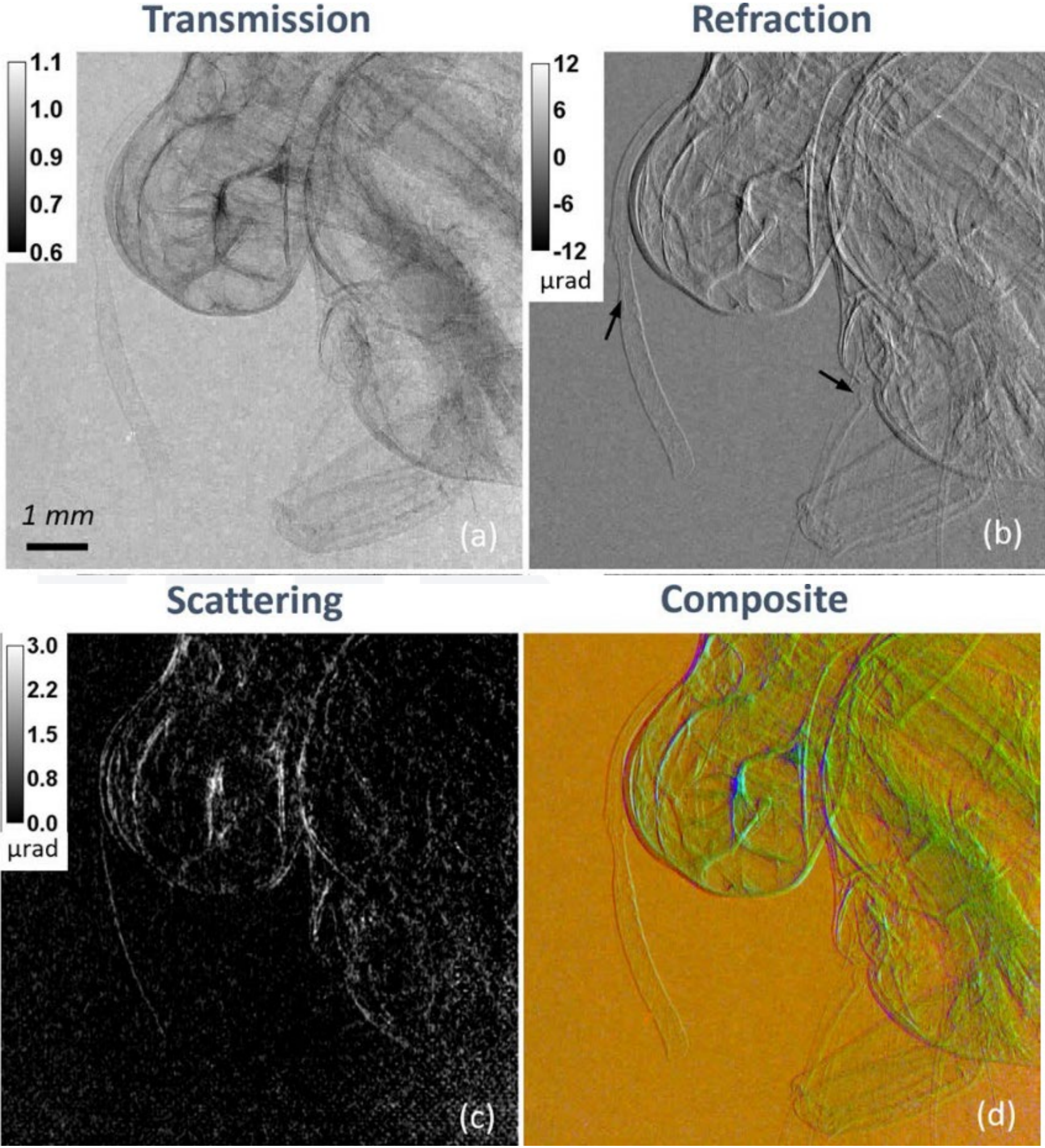
Both planar (2D) and tomography (3D)

M. Endrizzi *et al.*, Appl. Phys. Lett. 104, 024106 (2014)



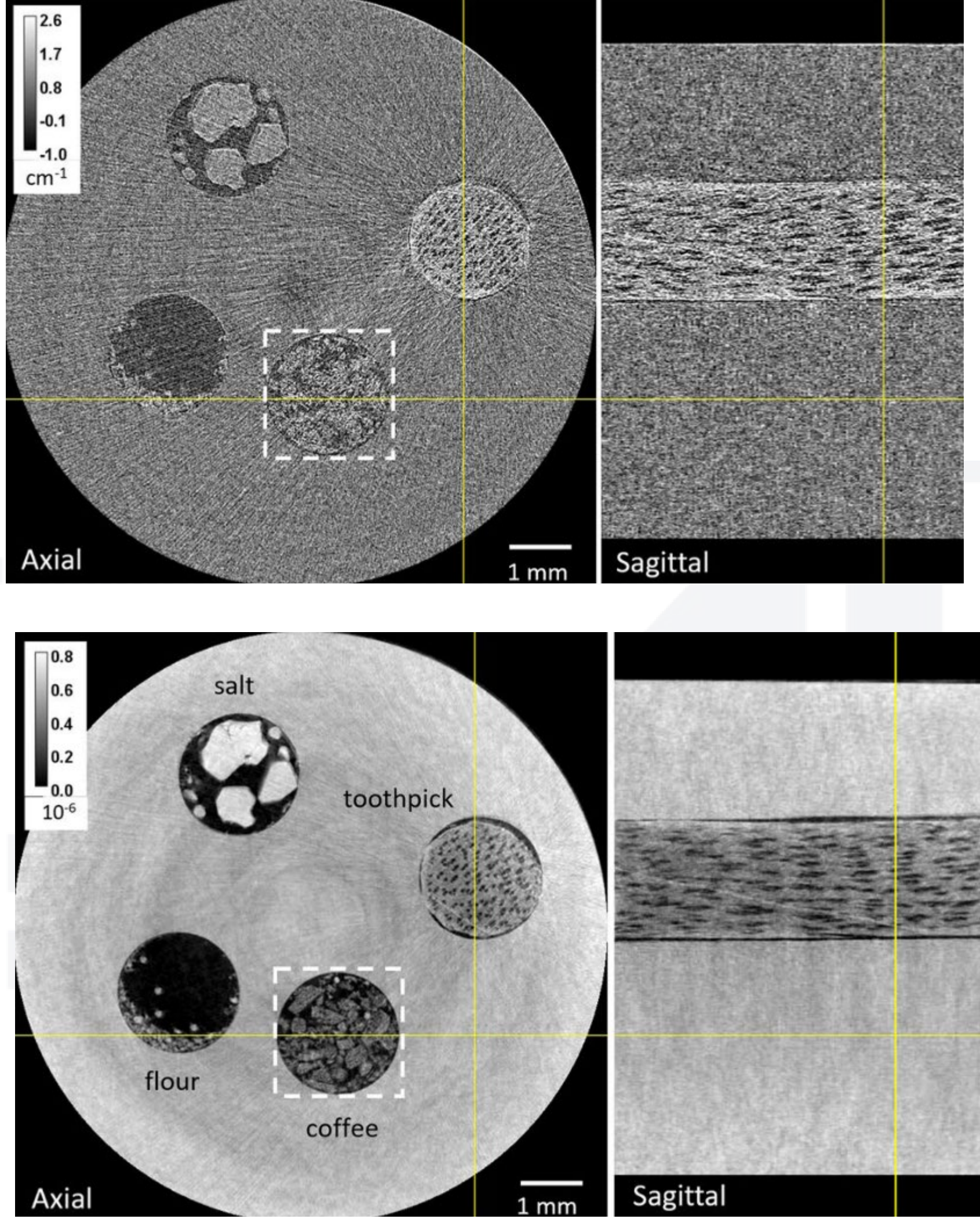
# PHASE-CONTRAST CAPABILITIES

Phase contrast radiography

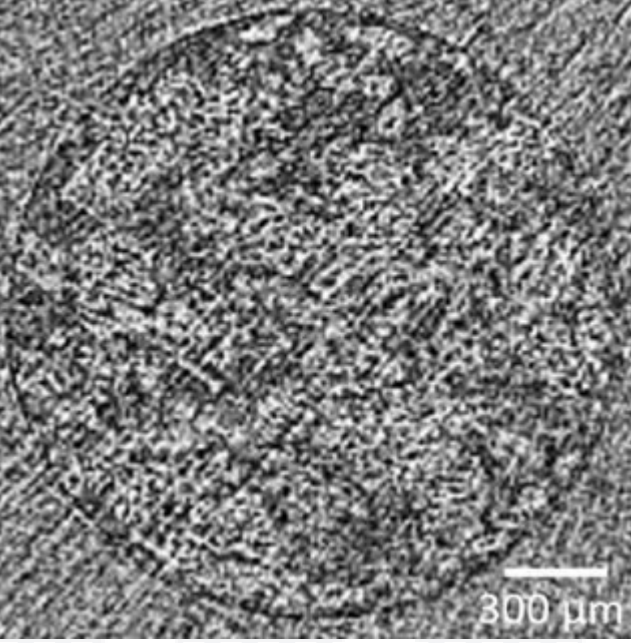


Brombal, Luca, et al. *Sci. Rep.* 13.1 (2023): 4206

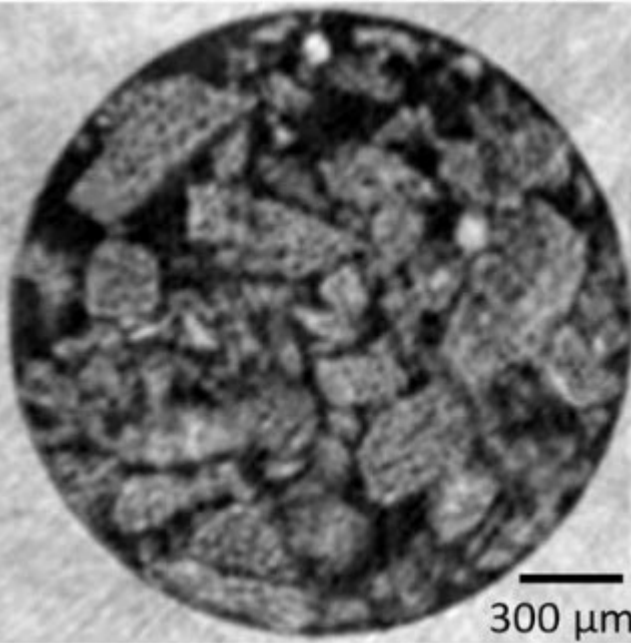
Phase contrast tomography



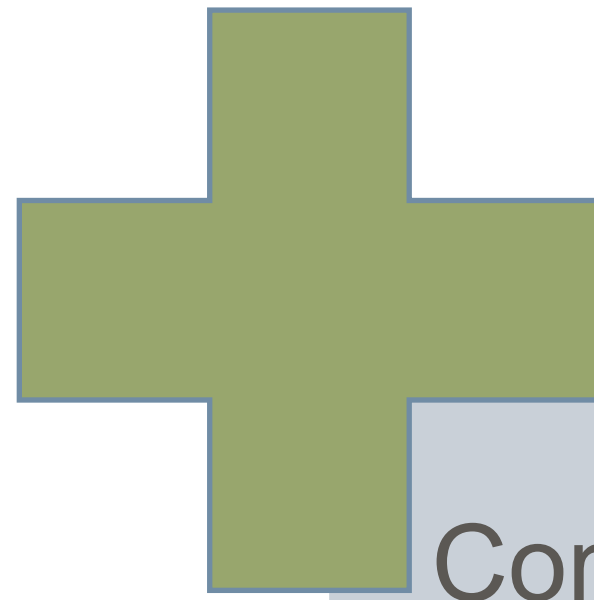
Attenuation



Phase



# EDGE ILLUMINATION PROS AND CONS



Compatible with low-coherence X-ray source

Works with polychromatic spectra

Flexible acquisition protocols (sensitivity vs speed, spatial resolution vs speed)

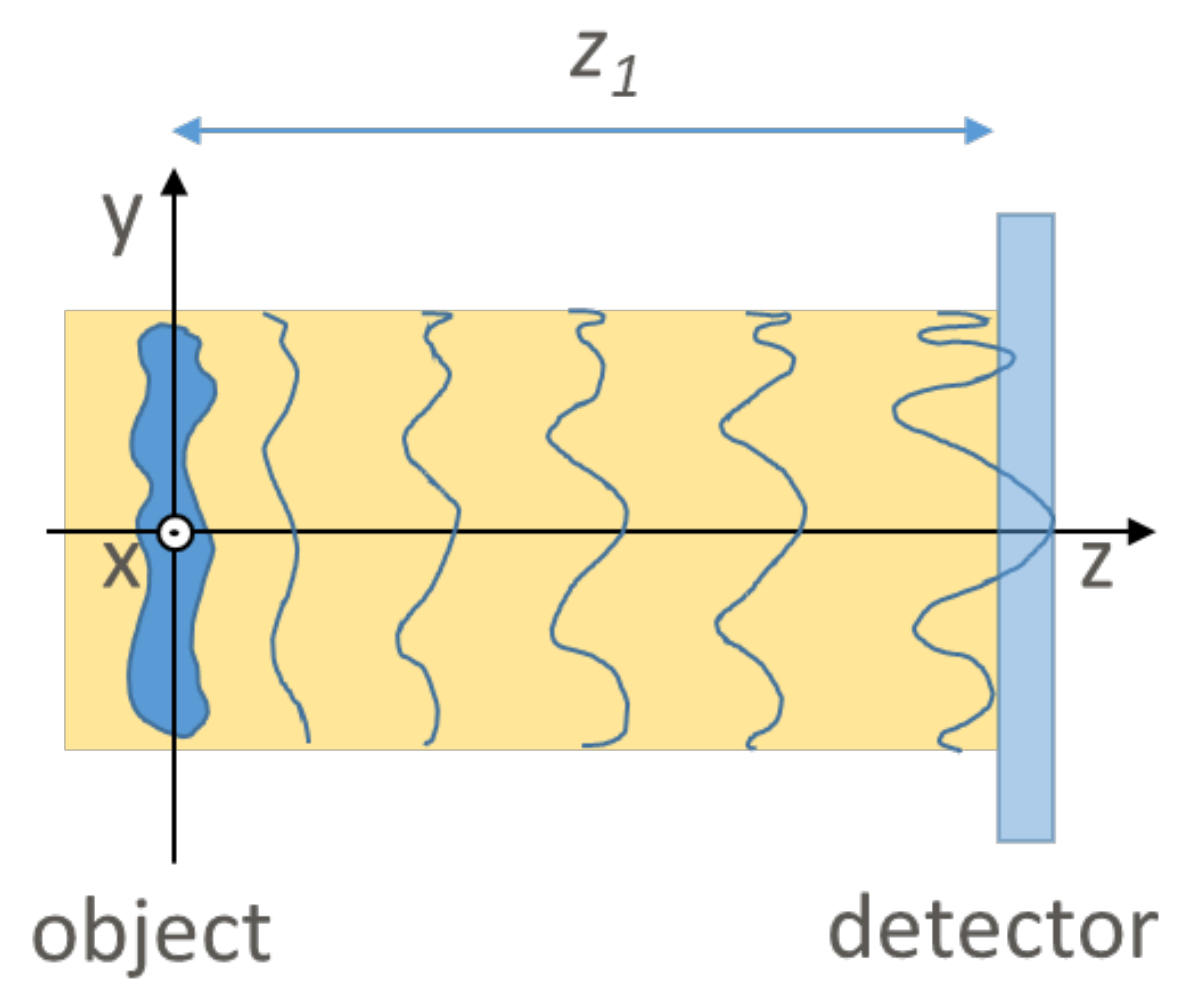


“Photon hungry” technique

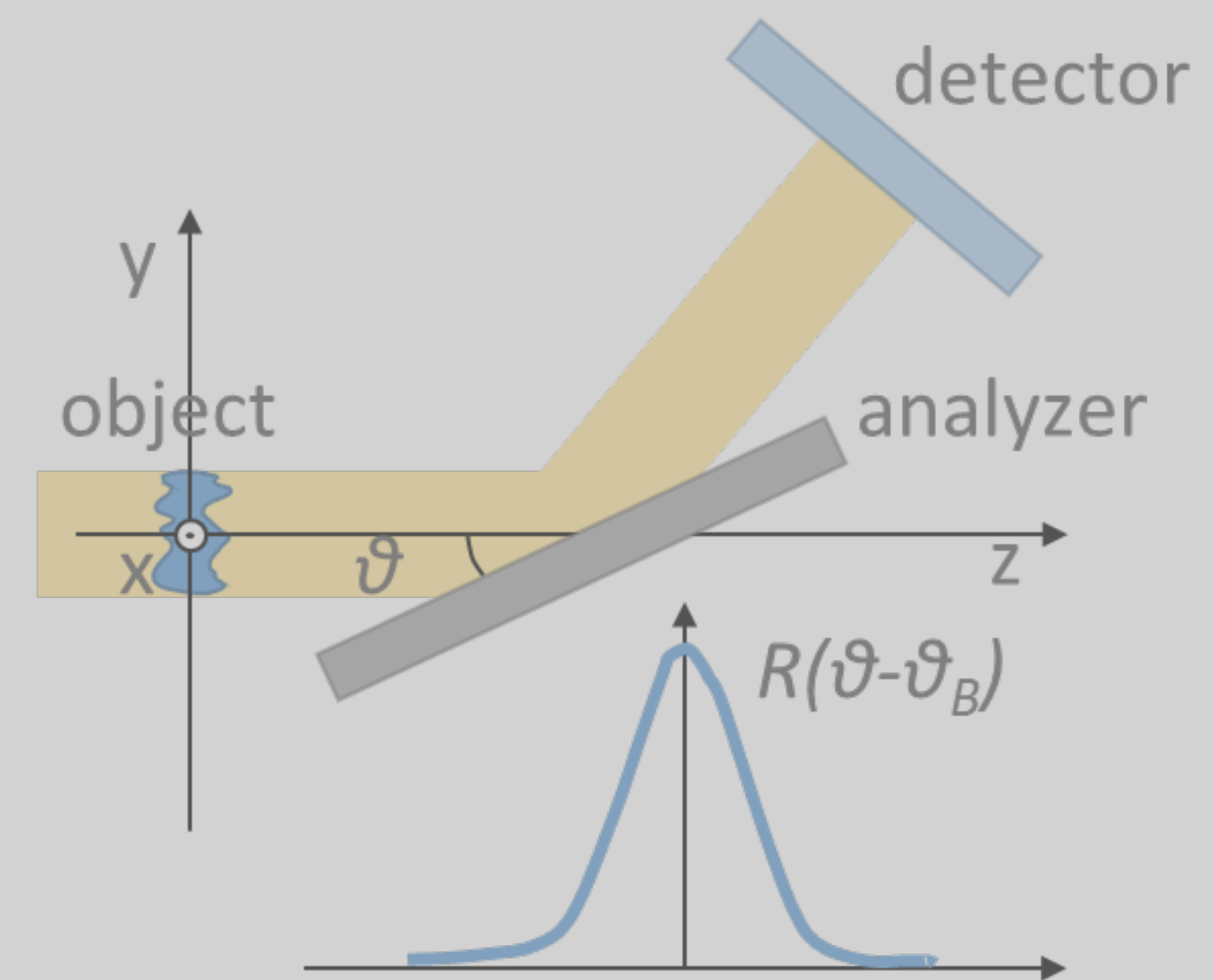
Alignment of masks is critical

Masks are “relatively” expensive

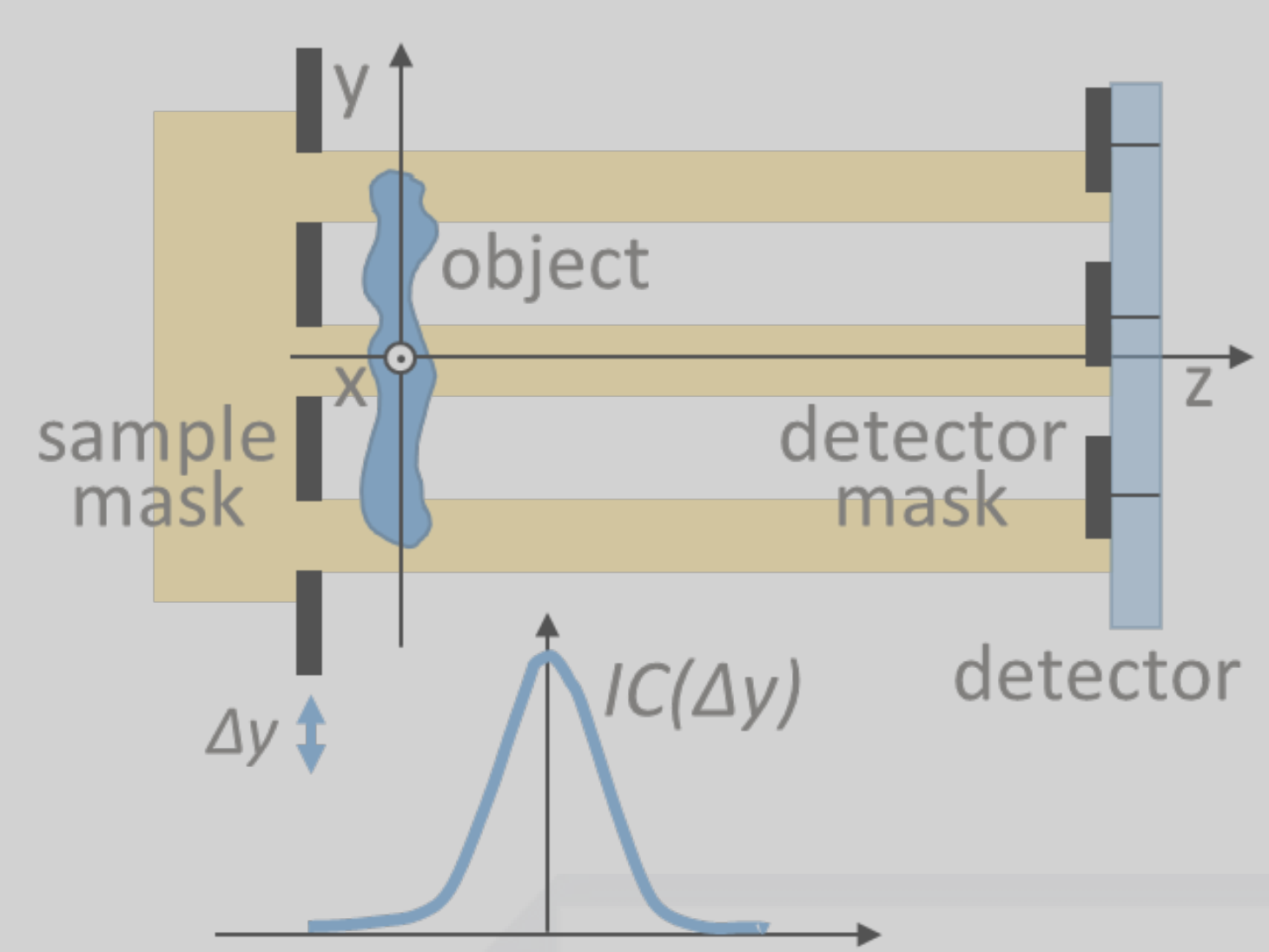
### PROPAGATION-BASED



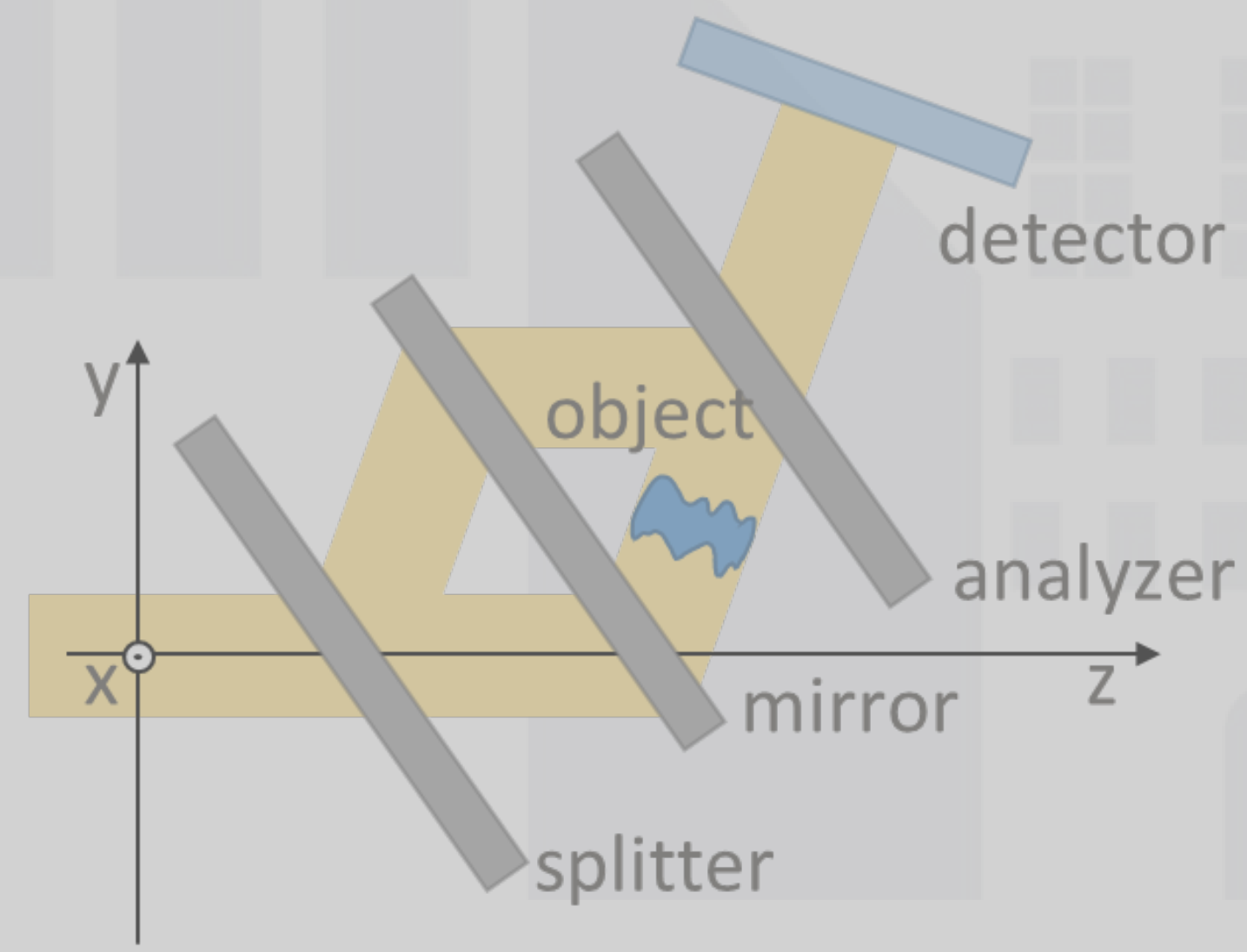
### ANALYZER-BASED



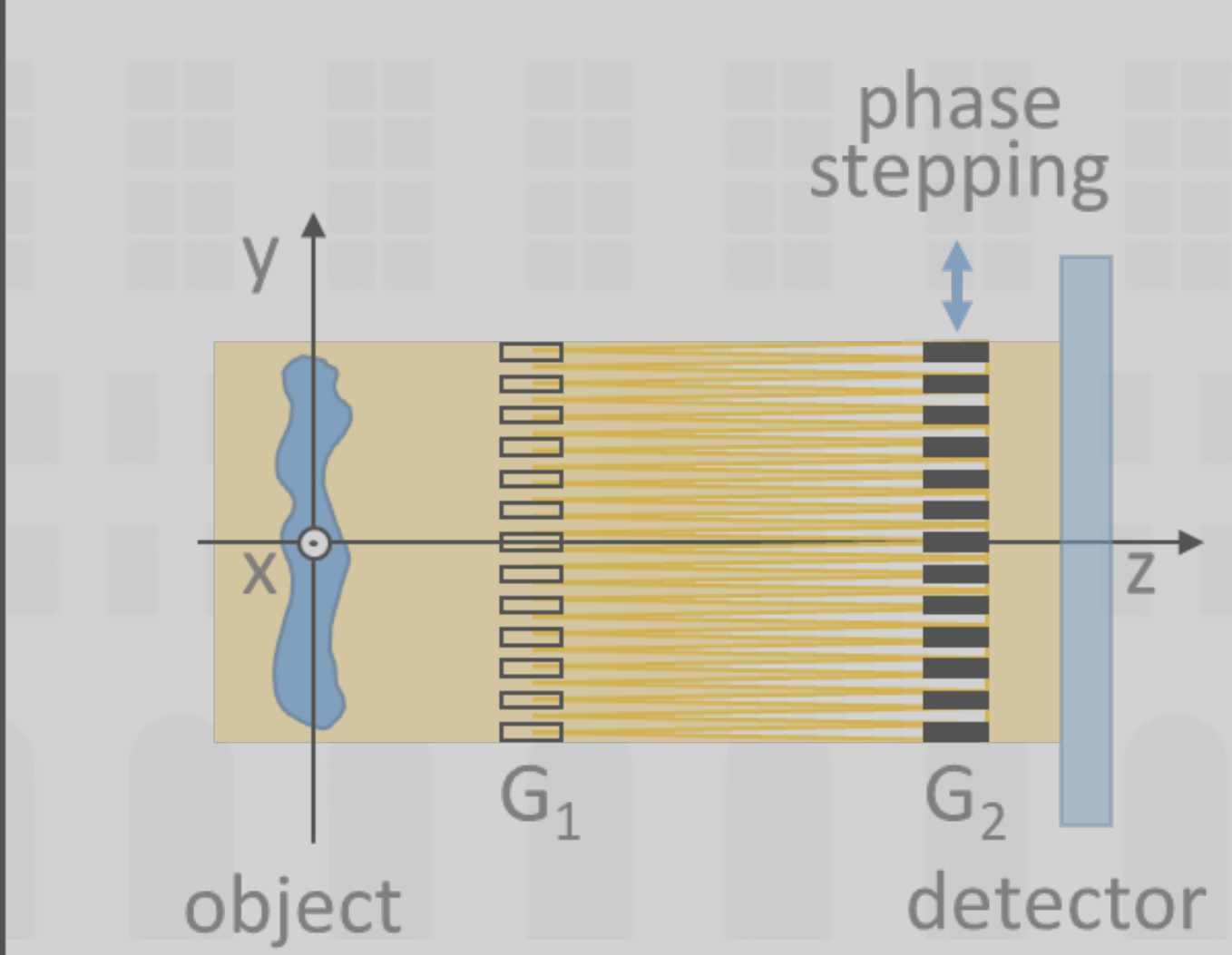
### EDGE ILLUMINATION



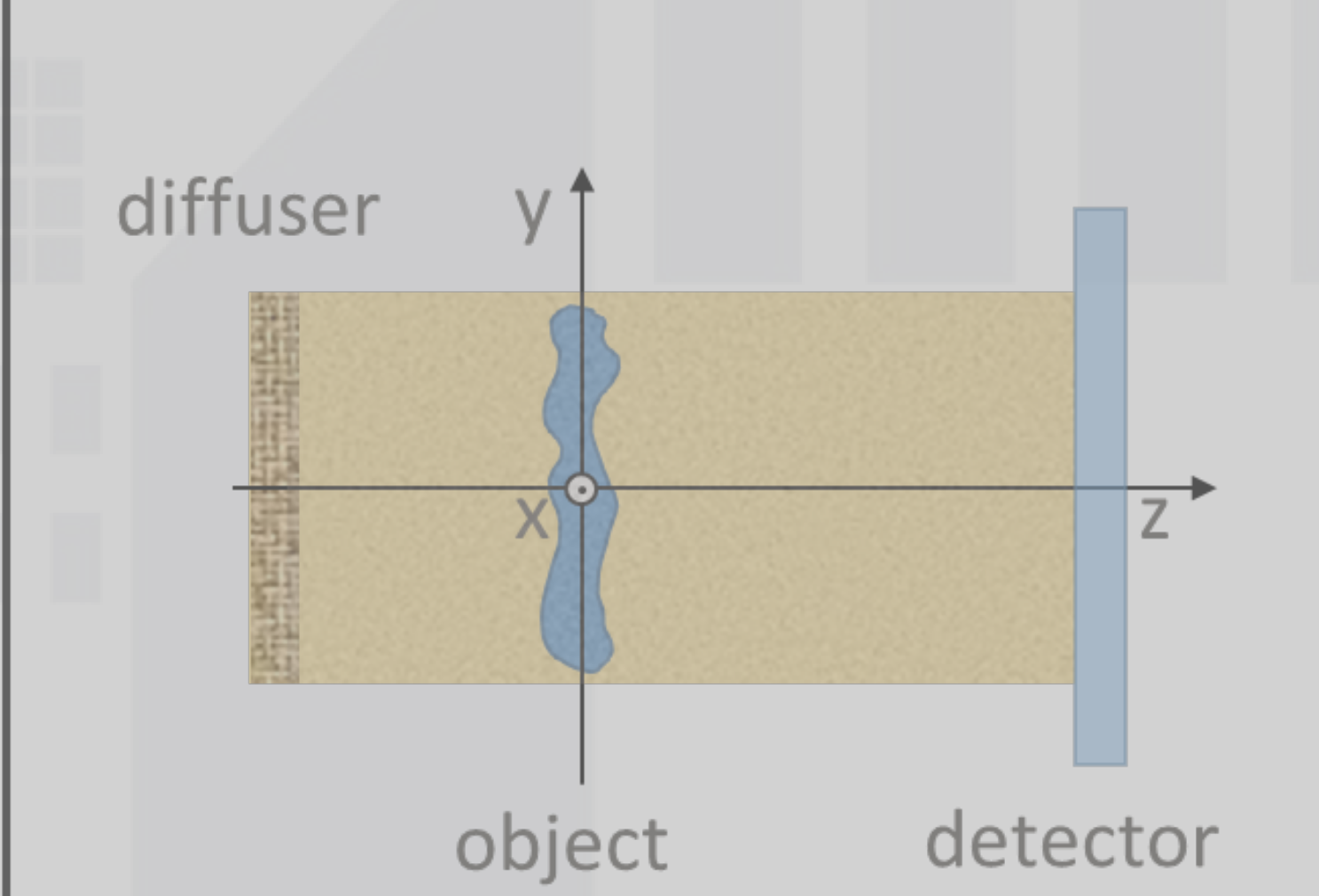
### INTERFEROMETRY



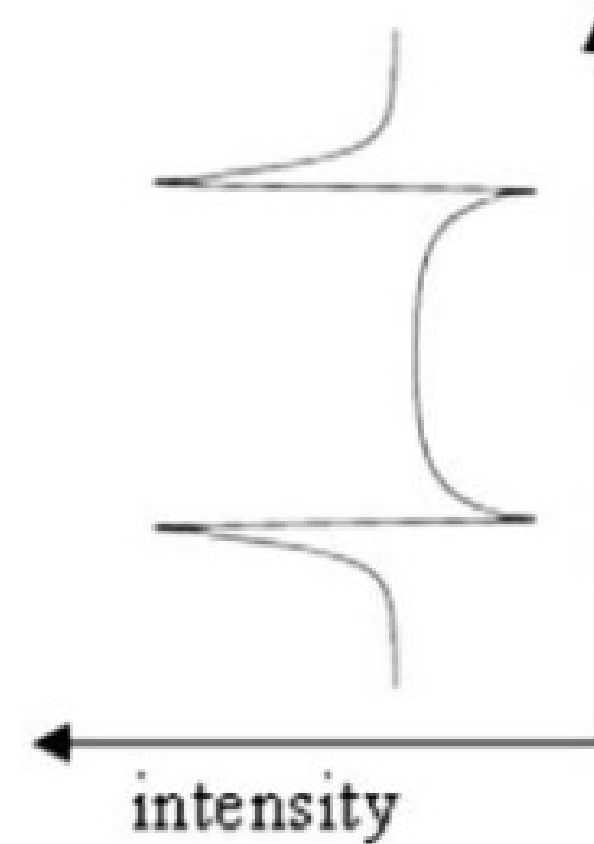
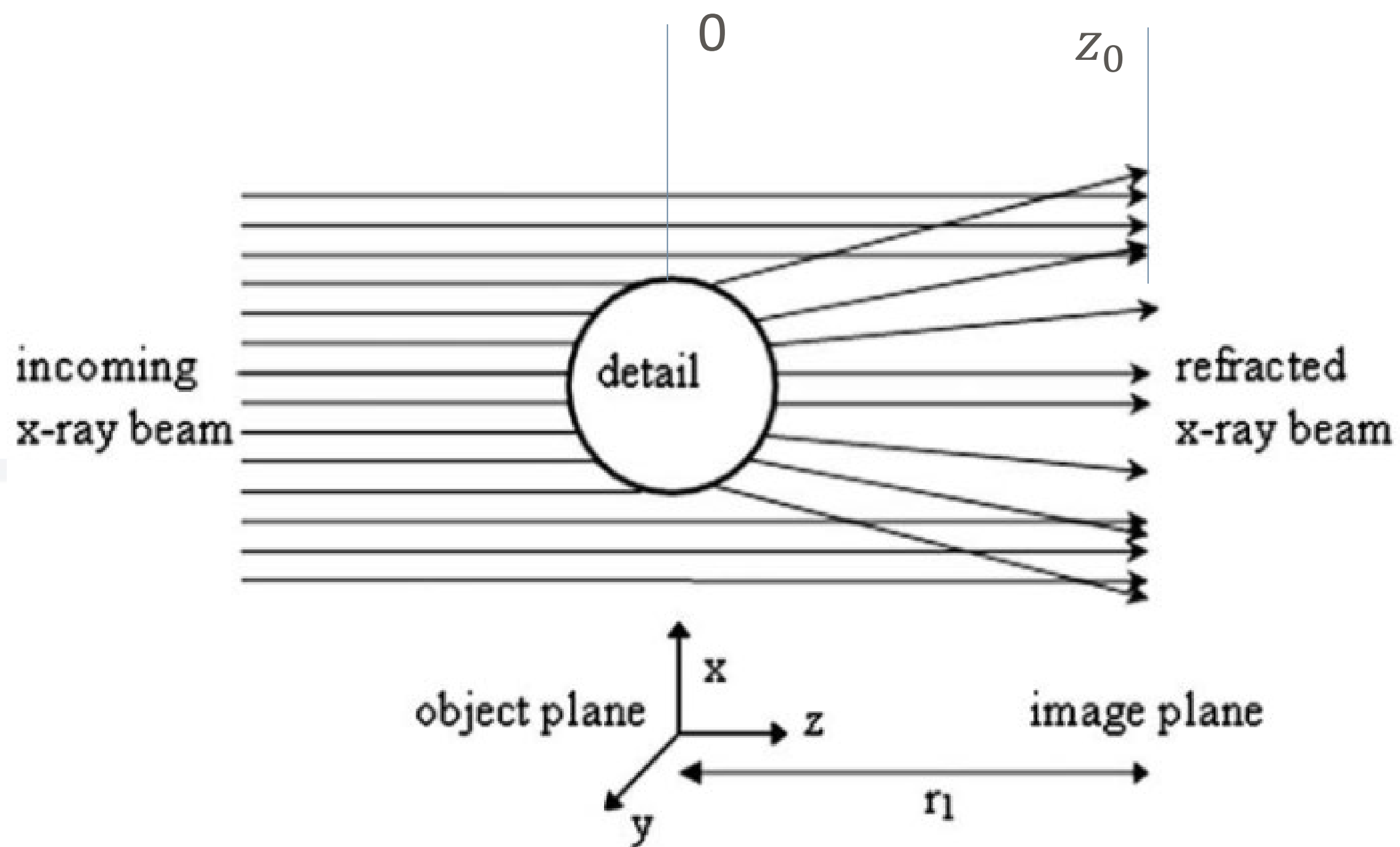
### GRATING INTERFEROMETRY



### SPECKLE IMAGING

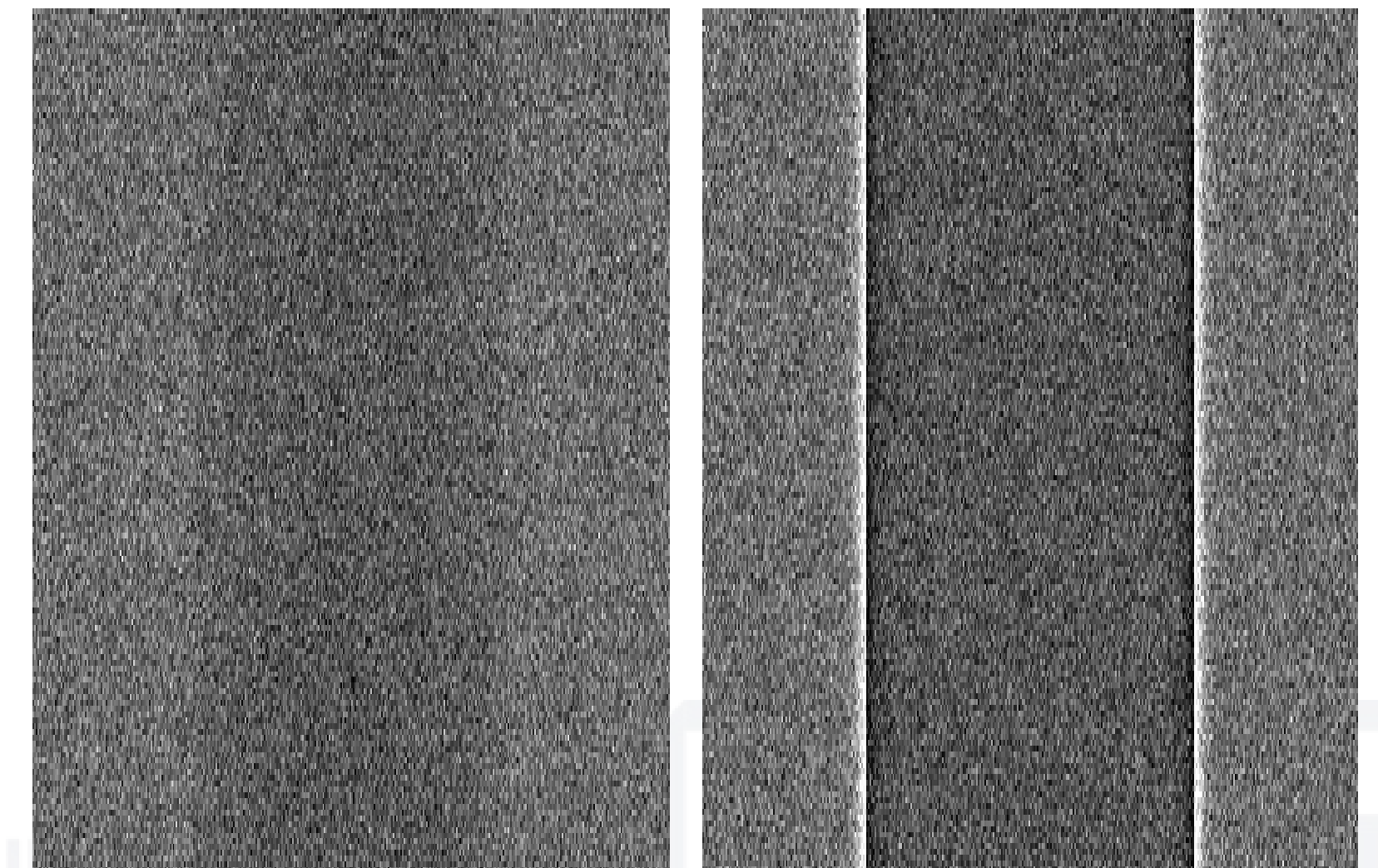


# PROPAGATION-BASED IMAGING - PBI



*No propagation*

*With propagation*



$$\psi_{out}(x, y; E) = \psi_{in}(E) e^{-k \int \beta(x, y, z; E) dz} e^{-ik \int \delta(x, y, z; E) dz}$$

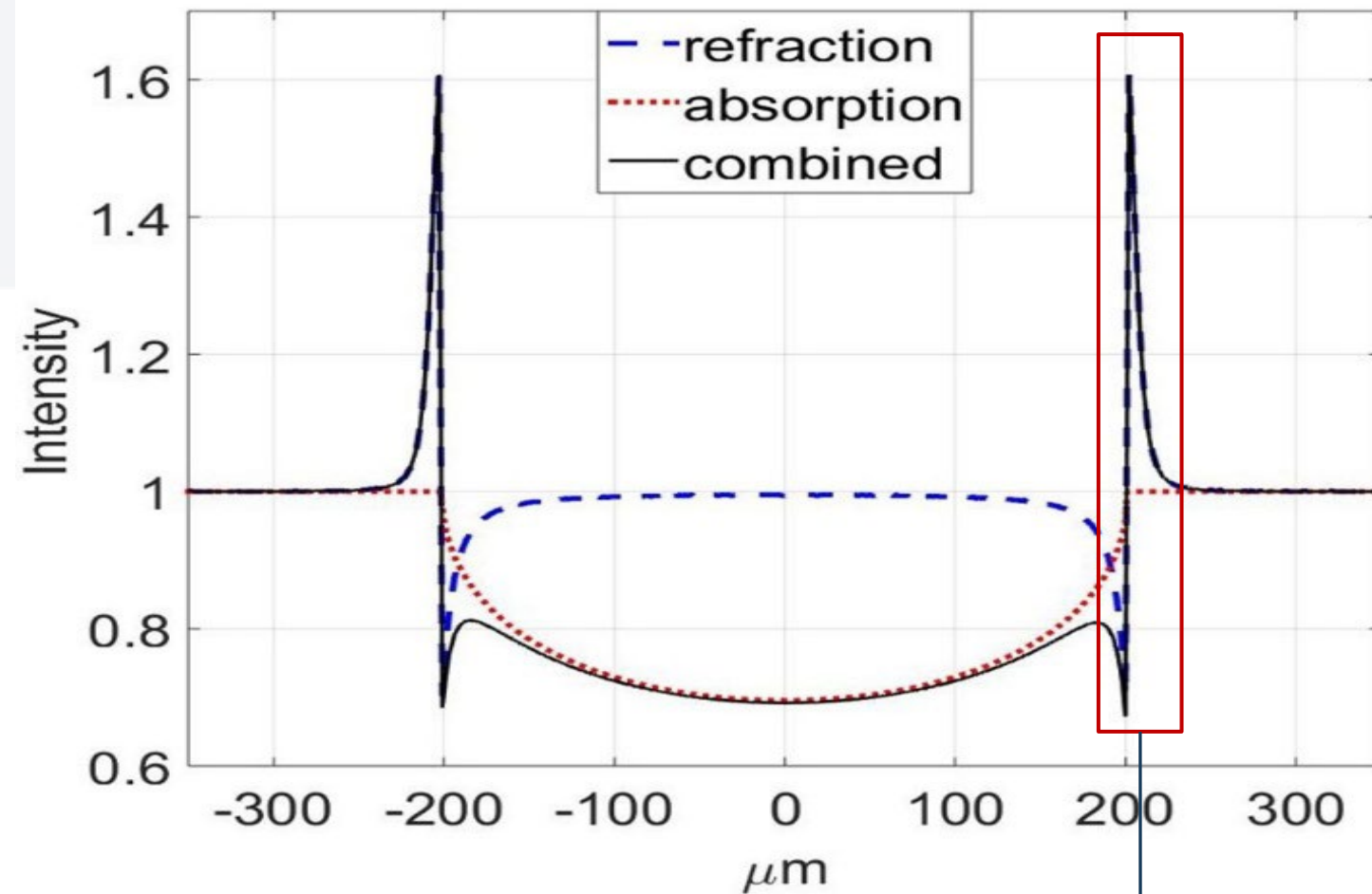
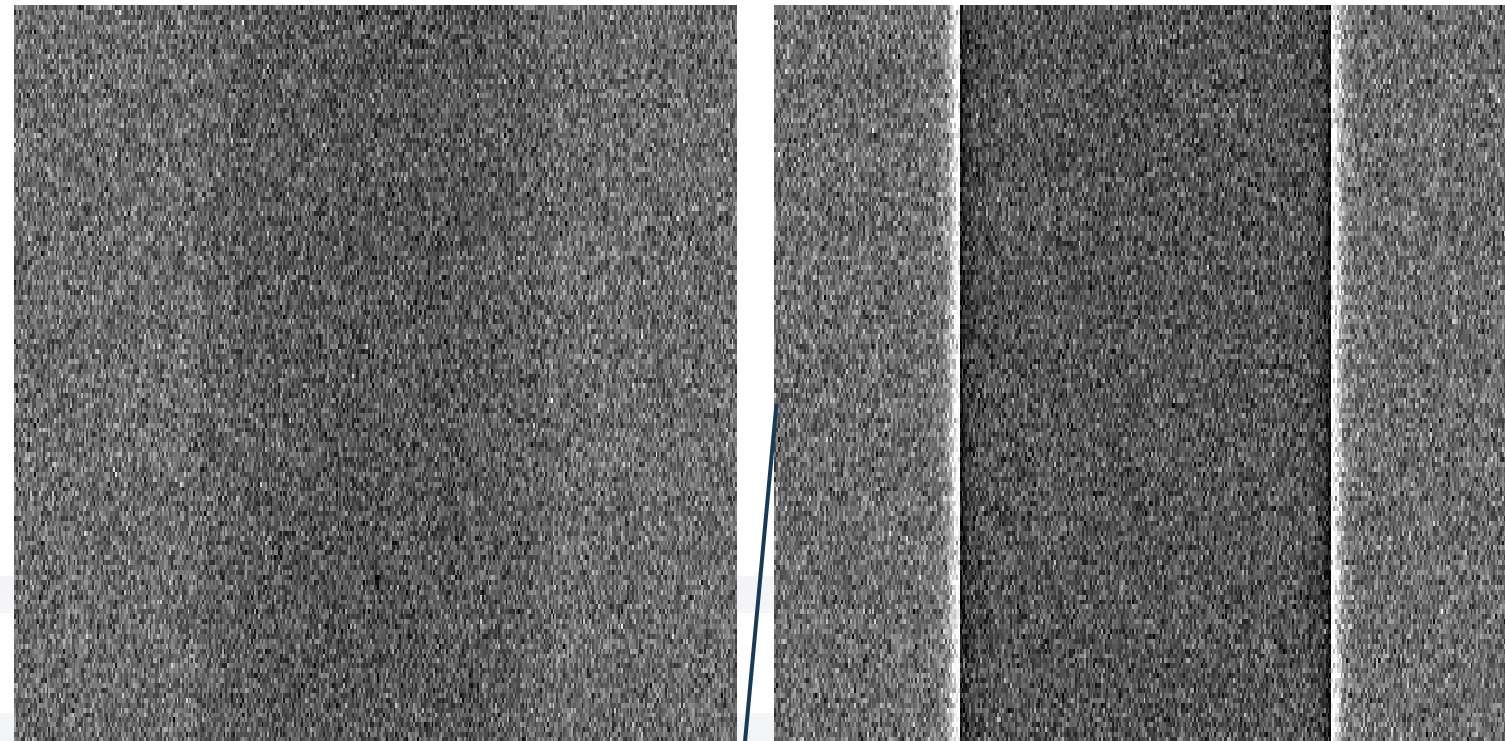
$$\Phi(x, y; E) = -k \int \delta(x, y, z; E) dz \quad \leftarrow \text{PHASE SHIFT}$$

Wilkins SW, et al. (1996) Nature 384: 335–338.

# X-RAY INTENSITY PROPAGATION (after some calculations...)

*No propagation*

*With propagation*



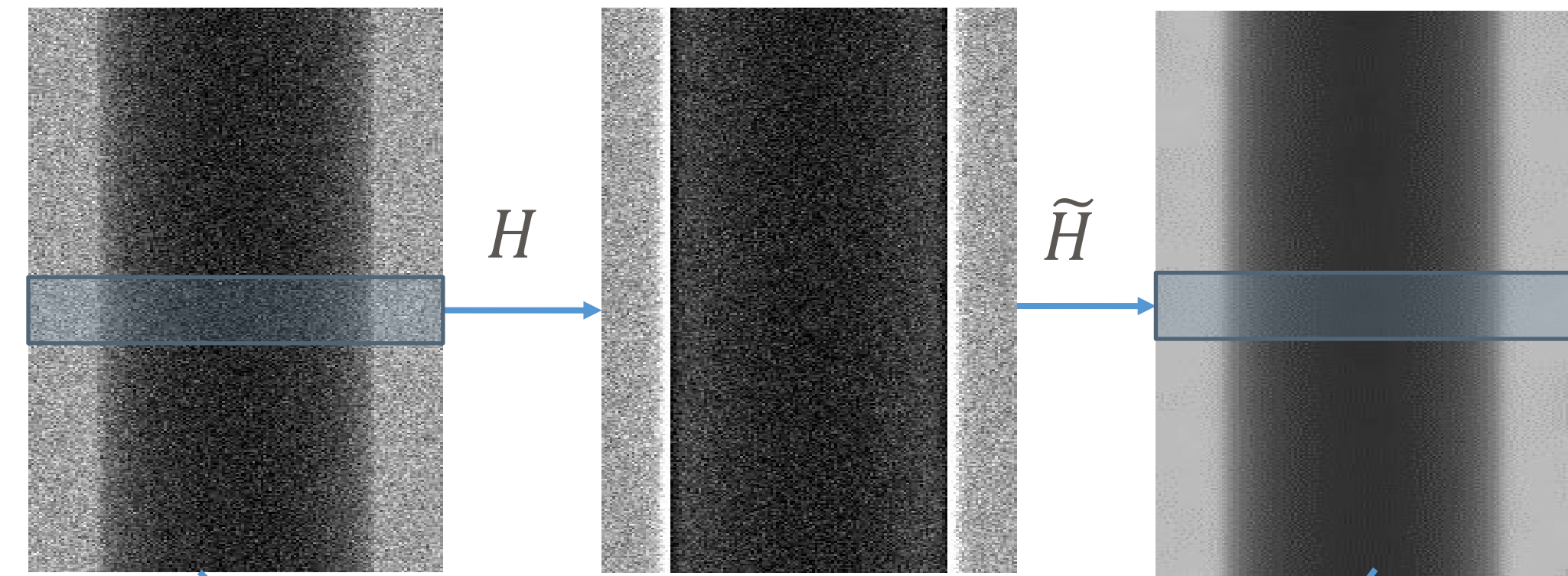
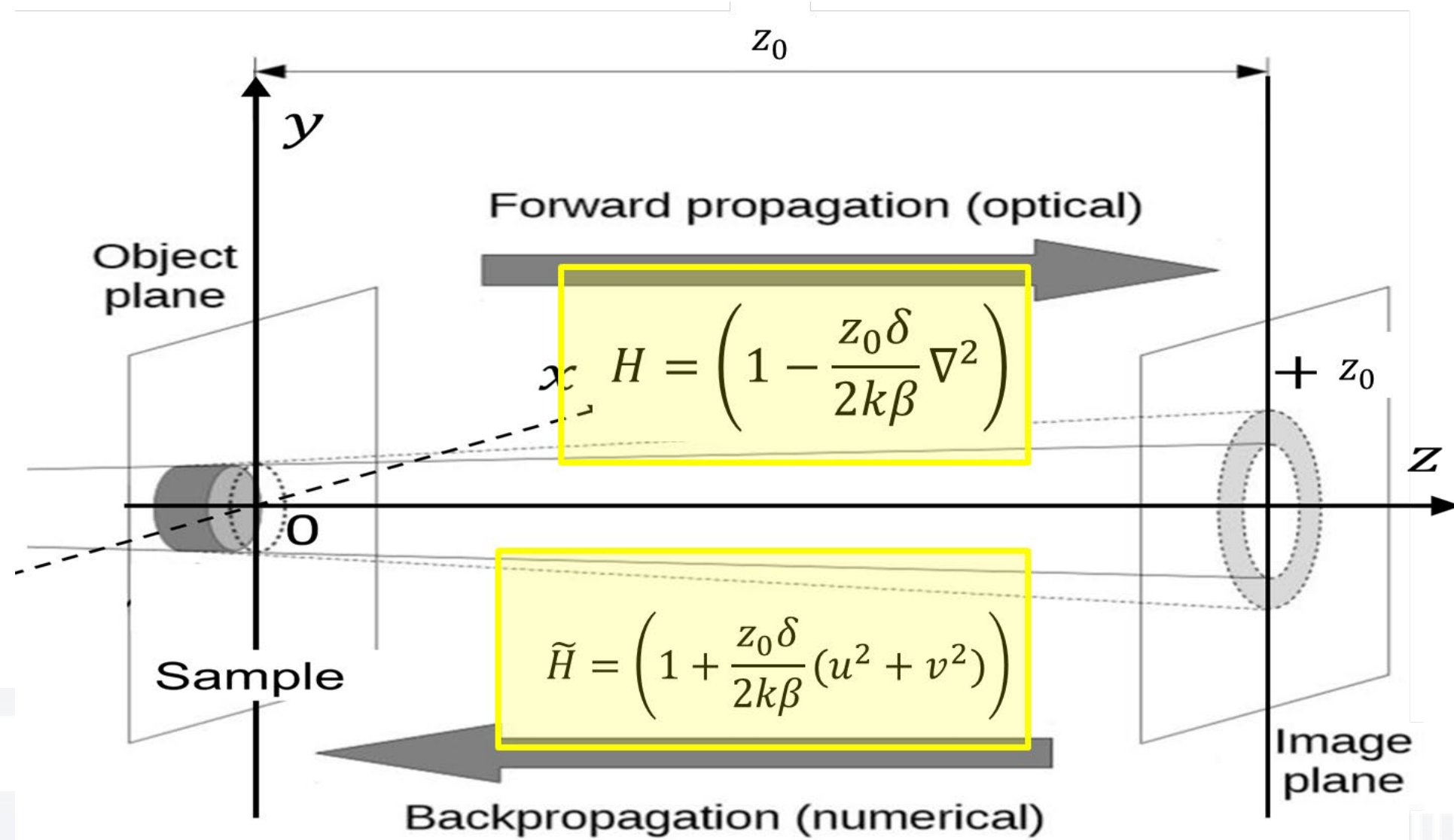
Edge enhancement

conventional  
attenuation image

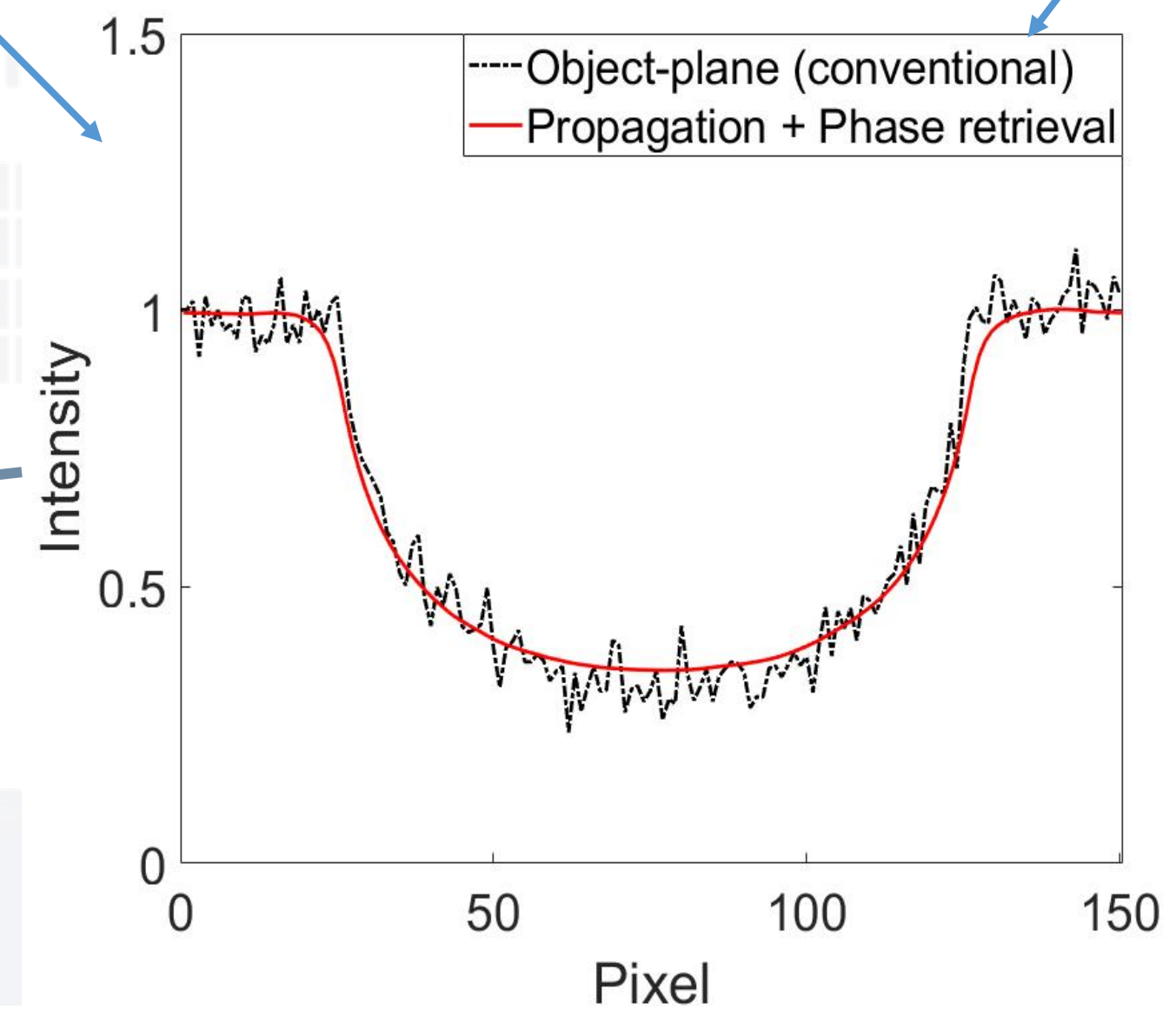
$$I(x, y; z = z_0) = I_0(x, y; z = 0) \left( 1 - \frac{z_0}{k} \nabla^2 \Phi(x, y) \right)$$

phase contrast term!  
 $\propto z_0$  (= propagation distance)  
 $\propto \nabla^2 \Phi(x, y)$

# PAGANIN'S PHASE-RETRIEVAL = "UNDOING PROPAGATION"



	Large area Contrast	Noise ( $\sigma$ )	SNR	Blur (resolution <sup>-1</sup> )
Propagation	=	=	=	↓
Phase retrieval	=	↓	↑	↑
Propagation + phase retrieval	=	↓	↑	=



❖ Gureyev et al., *J. Opt. Soc. Am. A* 34, 2251-2260 (2017)

❖ Brombal, Luca. *X-ray phase-contrast tomography: Underlying Physics and Developments for Breast Imaging*. Springer Nature, 2020.

# PROPAGATION-BASED PROS AND CONS



Easiest phase-contrast technique to implement

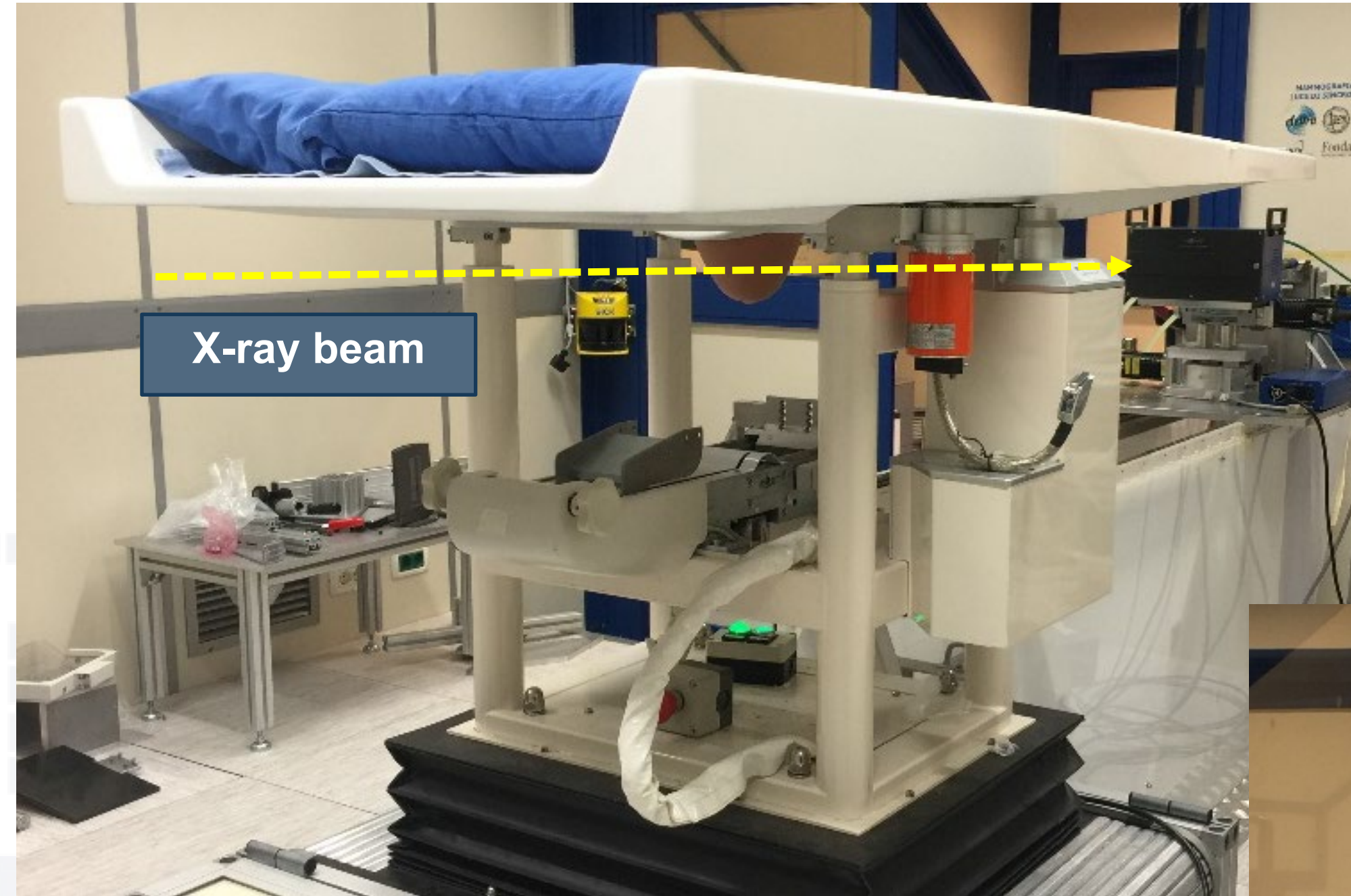
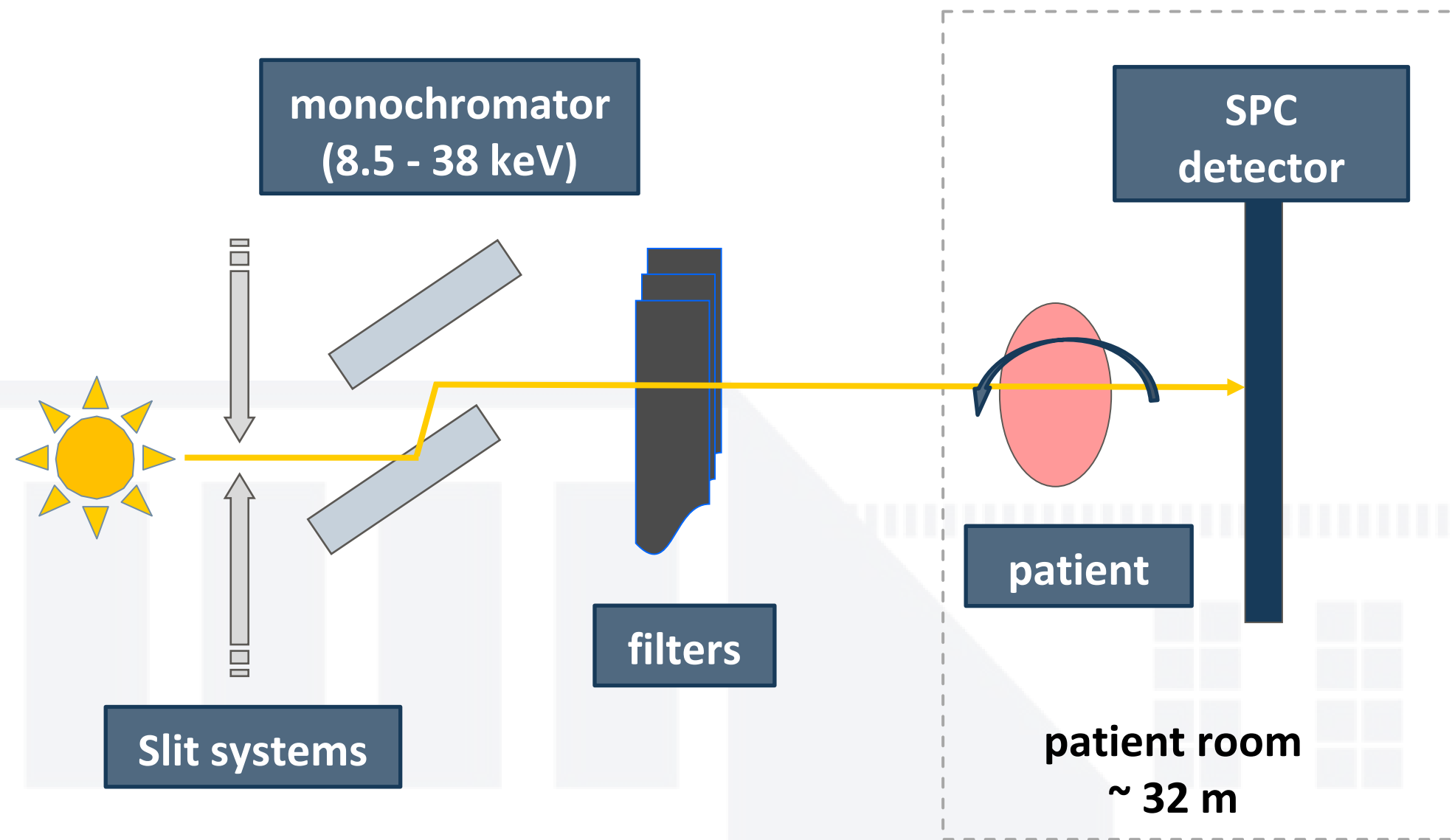
Widely used in research, robust data processing

A high-coherence X-ray source is required (low power or synchrotron)

It does not give access to different contrast channels separately (phase and attenuation are linked)

# PROPAGATION-BASED BREAST CT @ SYNCHROTRON

SYRMEP Beamline @ Elettra synchrotron

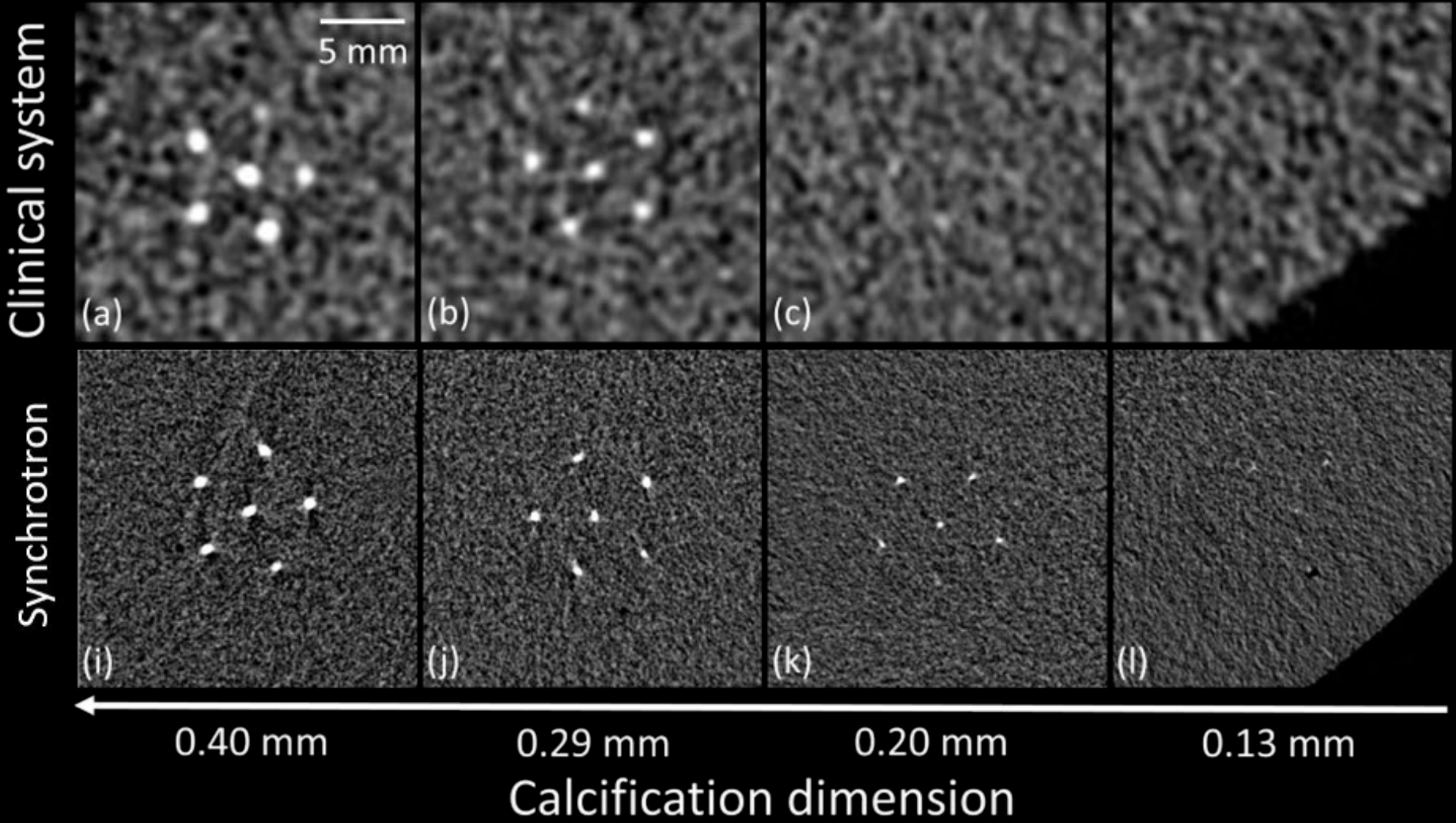
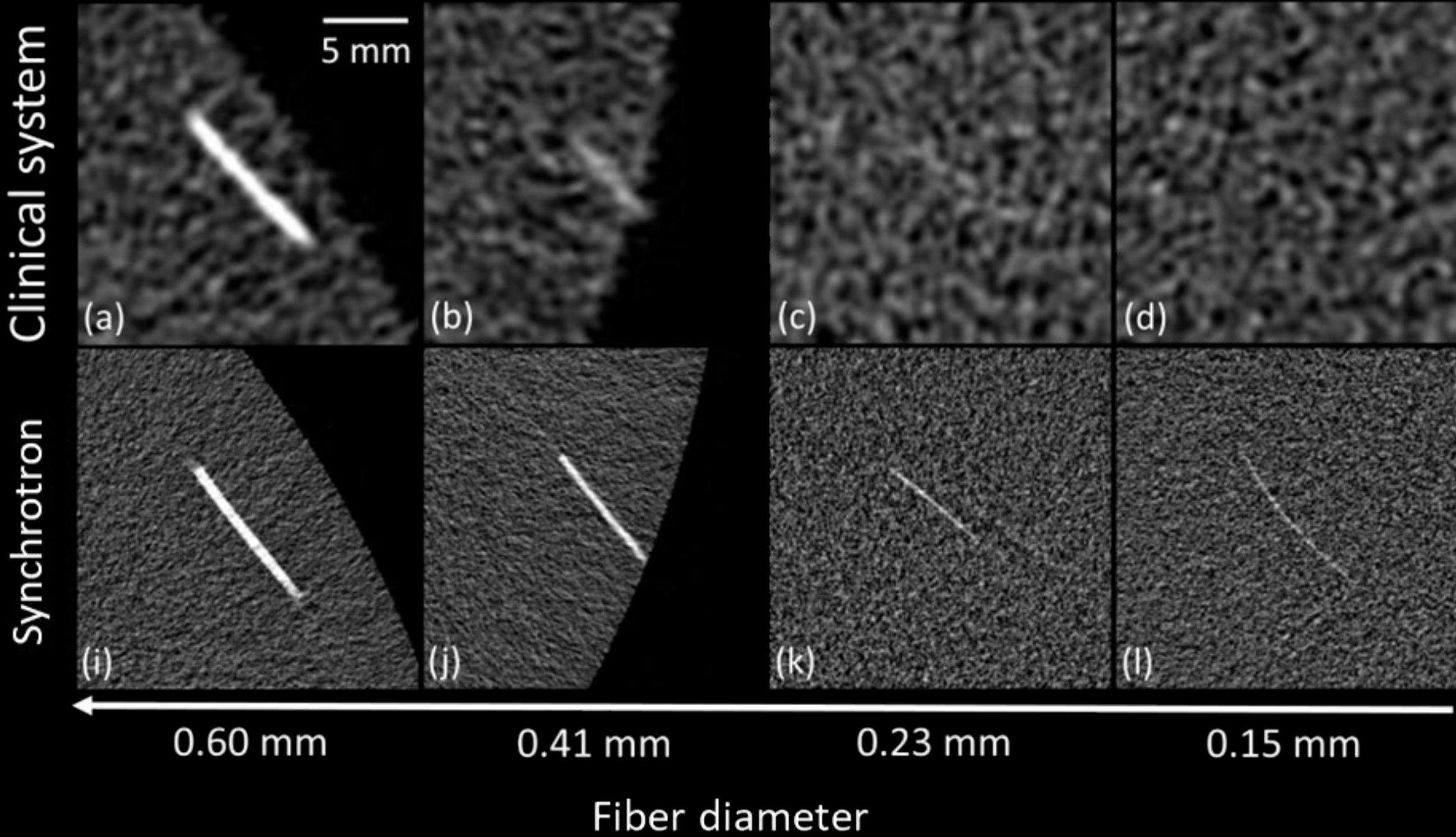


Elettra Sincrotrone Trieste



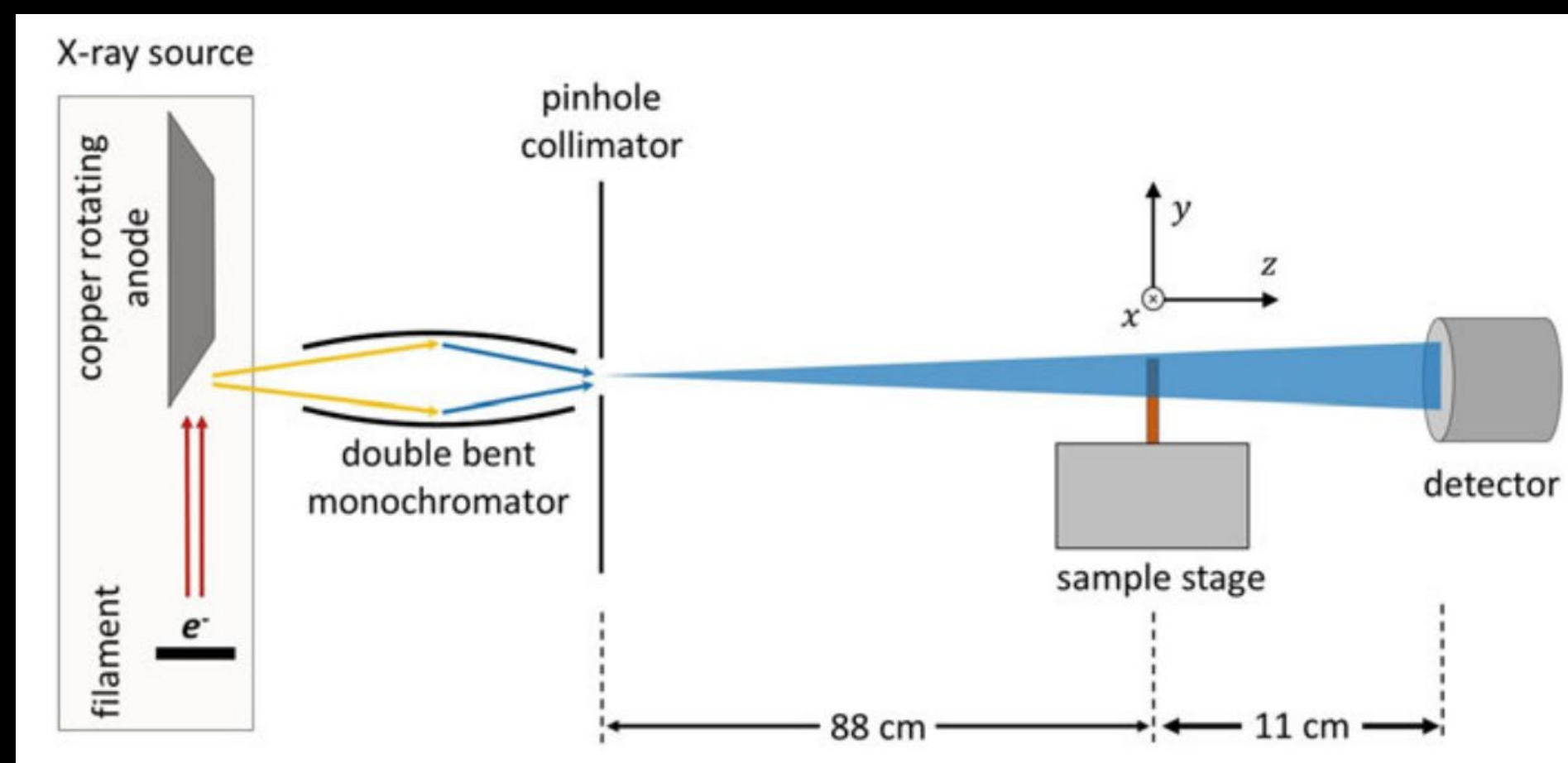
# PBI APPLICATIONS: CLINICAL VS PBI BCT

Clinically compatible radiation dose ~5 mGy

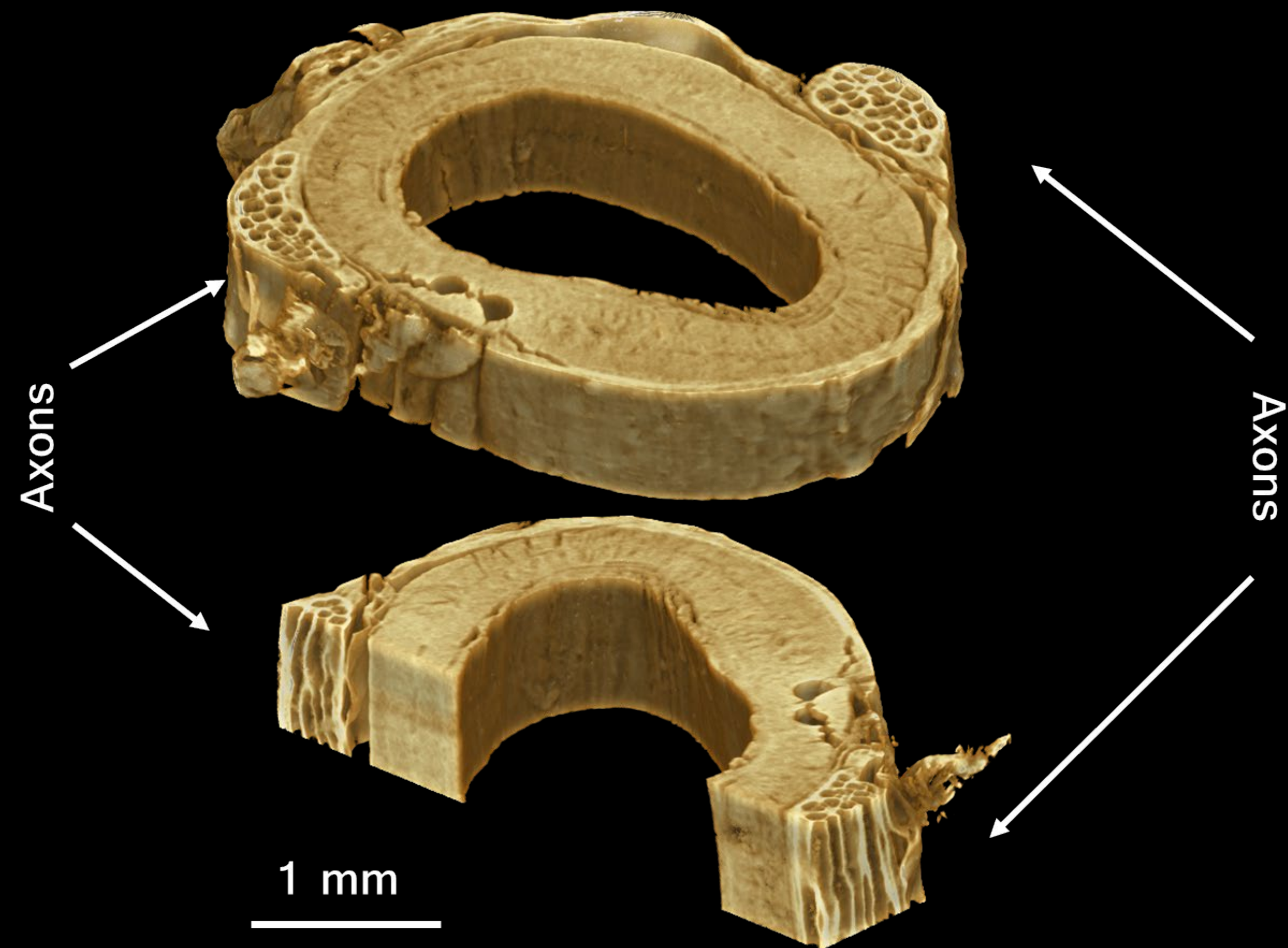


❖ Brombal, Luca, et al. "Image quality comparison between a phase-contrast synchrotron radiation breast CT and a clinical breast CT: a phantom based study." *Scientific reports* 9.1 (2019): 1-12.

# PBI APPLICATIONS: VIRTUAL HISTOLOGY IN THE LAB

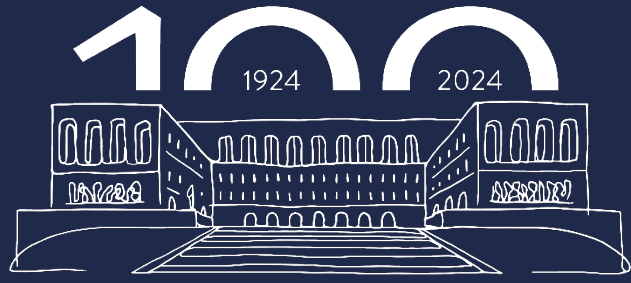


Piglet's esophagus



- i) Adventitia
- ii) Muscularis propria
- iii) Sub-mucosa
- iv) mucosa

*Brombal, L., et al. Physical Review Applied 11.3 (2019): 034004.*

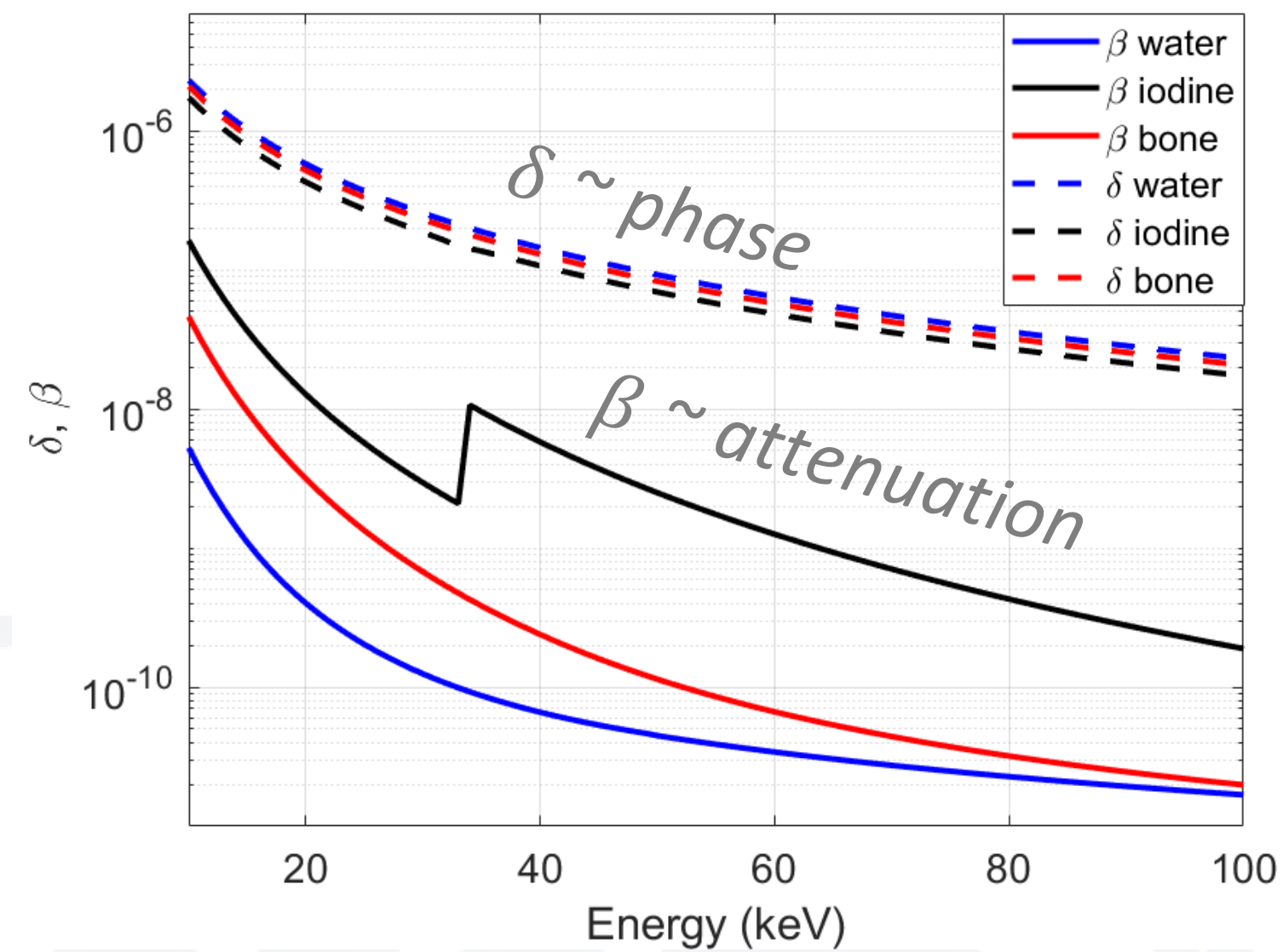


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# OUTLINE

- X-ray imaging fundamentals
- Spectral imaging
- Phase-contrast imaging
- **Spectral phase-contrast imaging**

# WHY SPECTRAL?



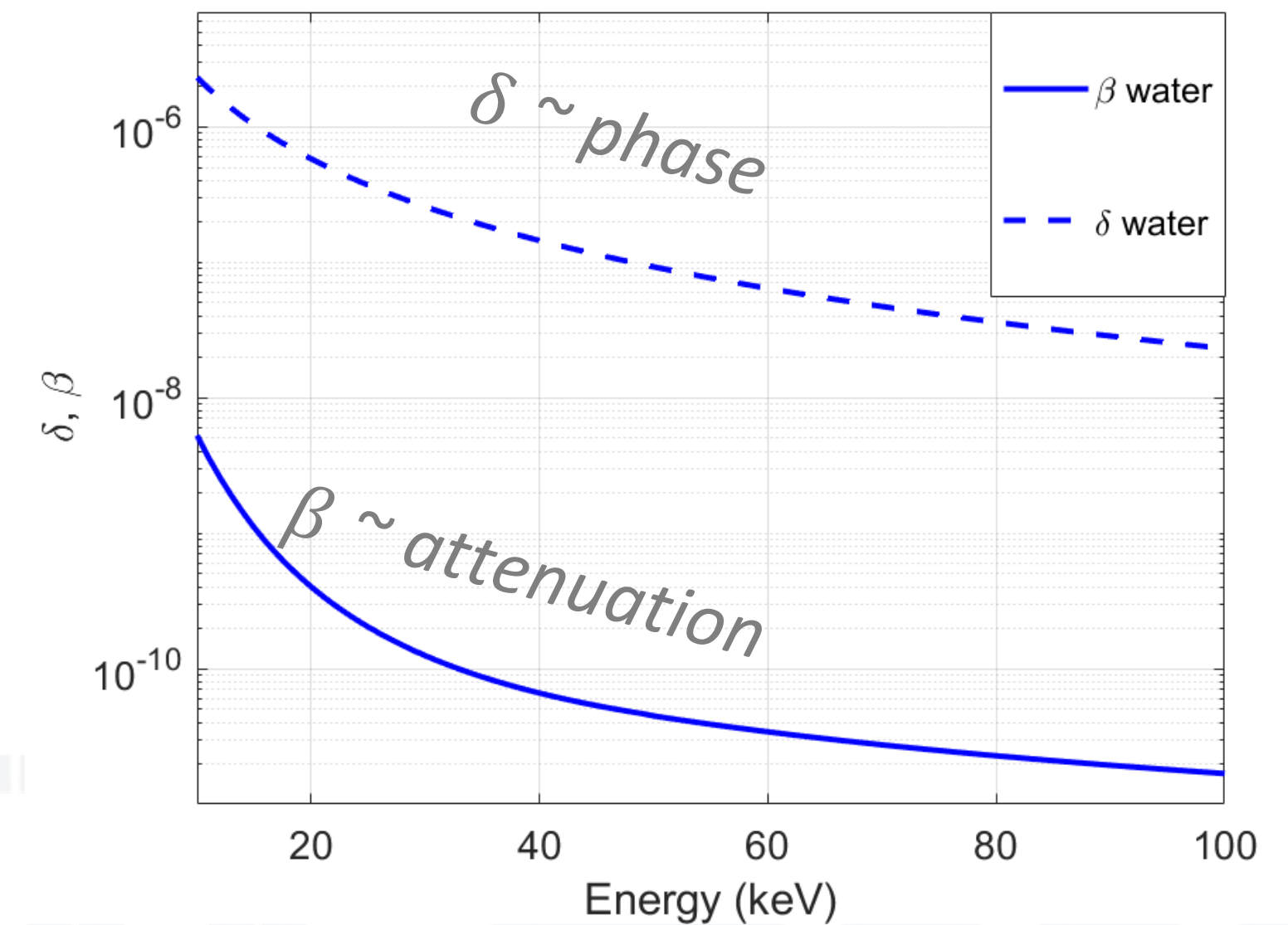
- Energy dependence of the attenuation coefficient is sensitive to chemical composition (i.e., difference in atomic number  $Z$ )

$$\mu(E) = \mu_{\text{photoelectric}}(E) + \mu_{\text{scattering}}(E) + \mu_{K_{\text{edge}}}(E)$$

$$\propto \frac{Z^4}{E^{3.5}} \quad \propto Z \quad \neq 0 \text{ at } K_{\text{edge}}$$

**to separate and quantify chemical elements!**

# WHY PHASE-CONTRAST?

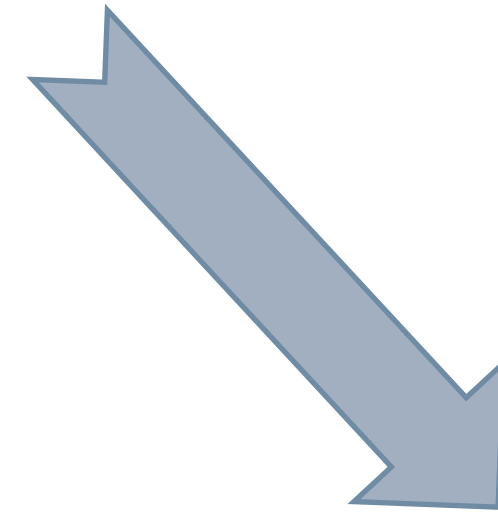


- For light materials and energies in the range 10 – 100 keV phase effects are much larger than attenuation

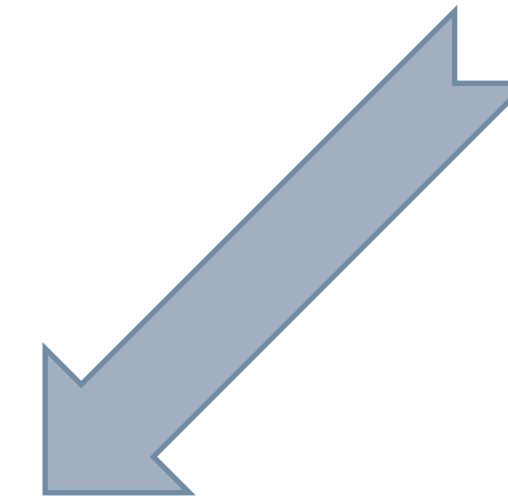
$$\frac{\delta}{\beta} \sim 10^3$$

**to increase visibility of soft tissues!**

# SPECTRAL IMAGING



# PHASE-CONTRAST IMAGING

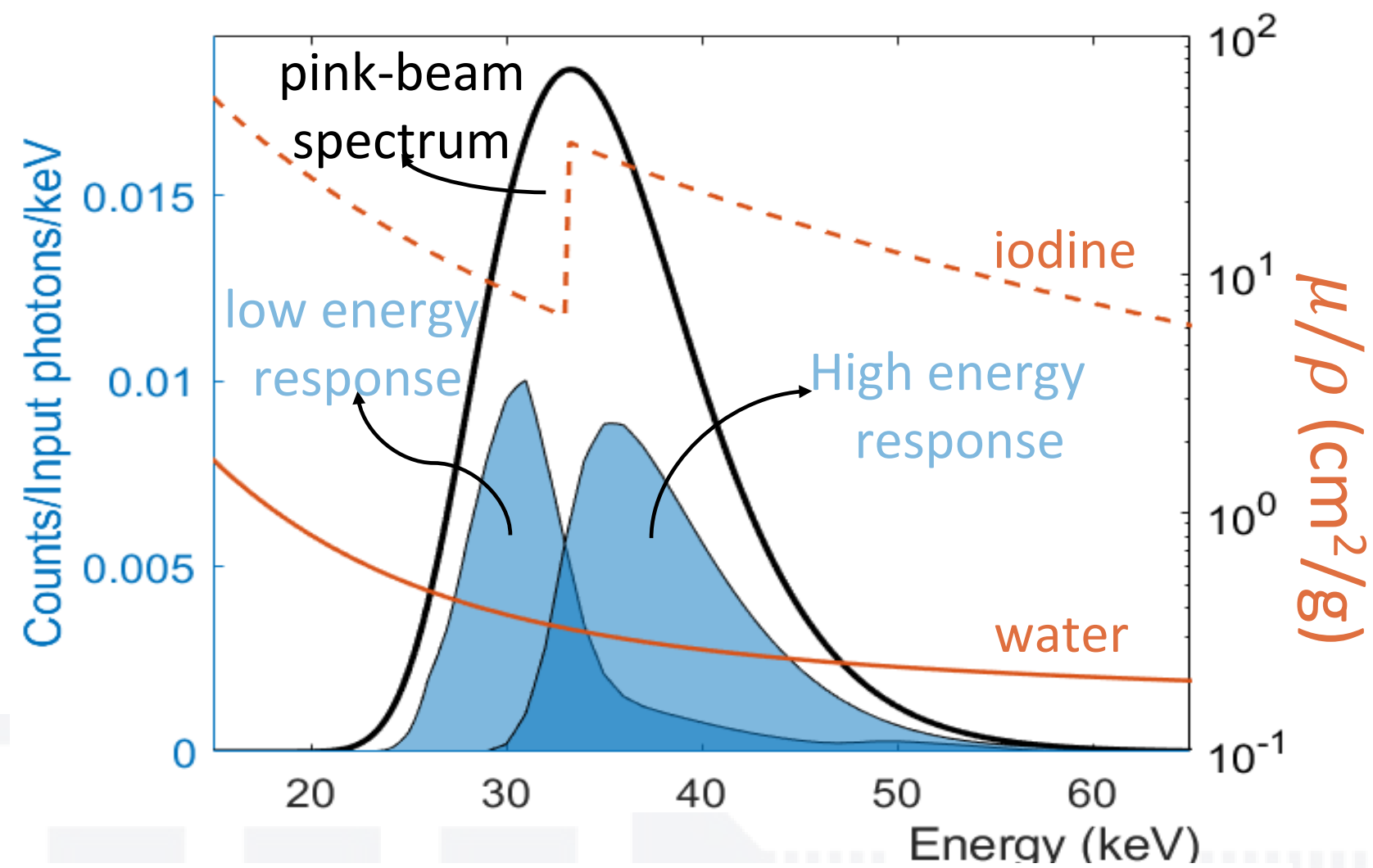


## SPECTRAL PHASE-CONTRAST IMAGING



- ✓ BOTH SOFT TISSUE VISIBILITY AND MATERIAL QUANTIFICATION
- ✓ ROBUST MATERIAL DECOMPOSITION THANKS TO THE LOW-NOISE PHASE-CHANNEL
- ✓ DECOMPOSITION OF 1 ADDITIONAL MATERIAL BY ADDING PHASE CHANNEL INTO DECOMPOSITION MATRIX

# SPECTRAL PHASE-CONTRAST MATERIAL DECOMPOSITION

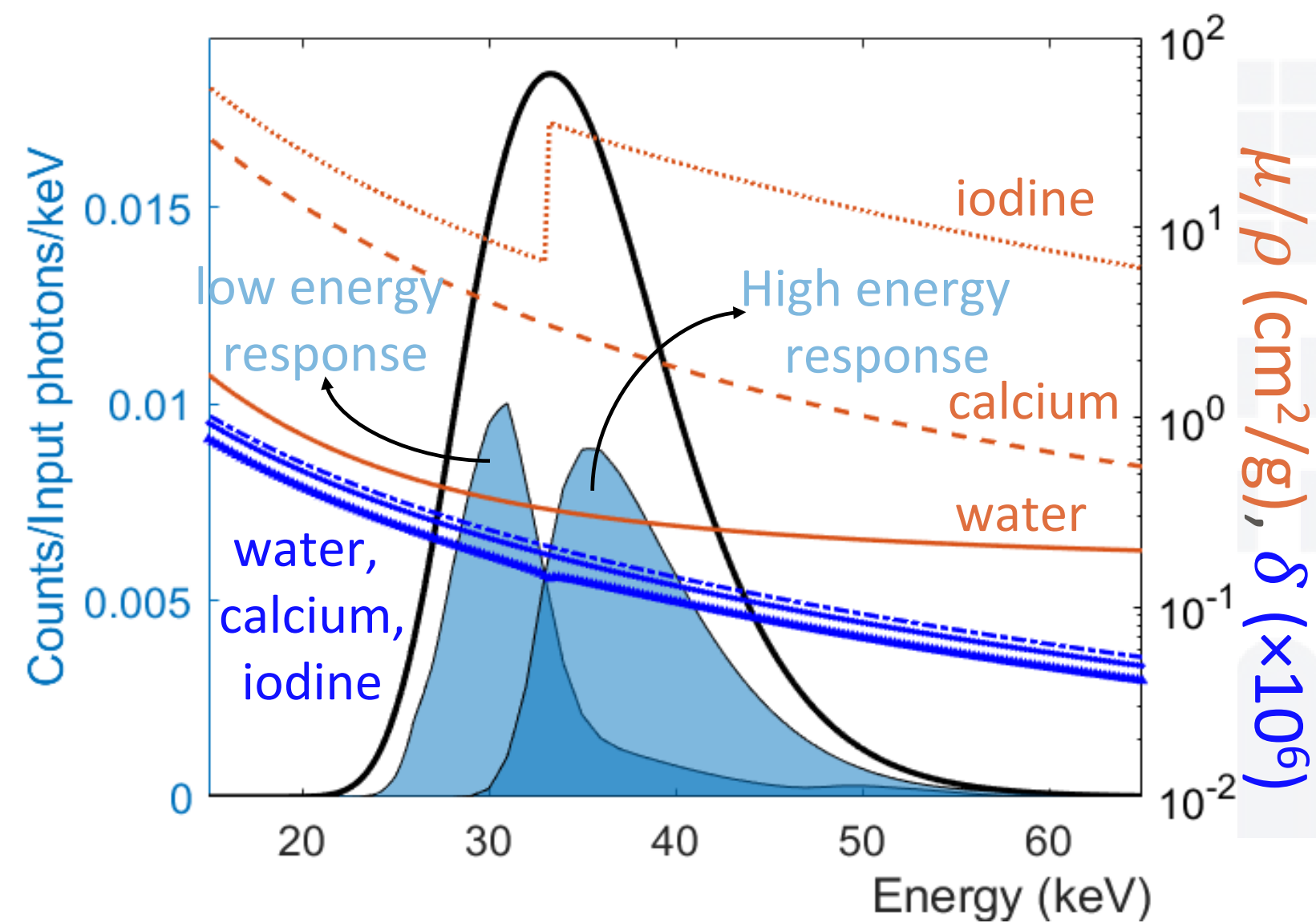


$$\mu_{low} = \frac{\mu}{\rho} \Big|_{low}^{water} \rho^{water} + \frac{\mu}{\rho} \Big|_{low}^{iodine} \rho^{iodine}$$

$$\mu_{high} = \frac{\mu}{\rho} \Big|_{high}^{water} \rho^{water} + \frac{\mu}{\rho} \Big|_{high}^{iodine} \rho^{iodine}$$

What if we want a third material?

Add the phase ( $\delta$ ) channel!



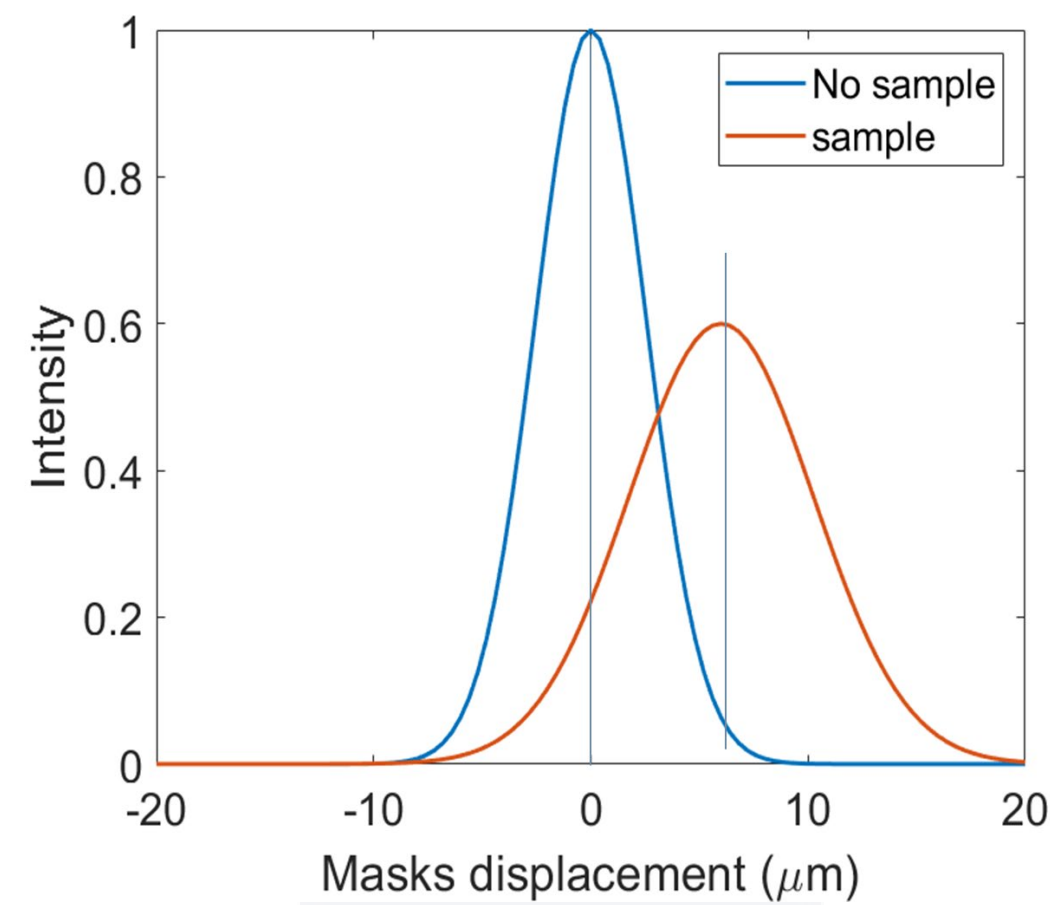
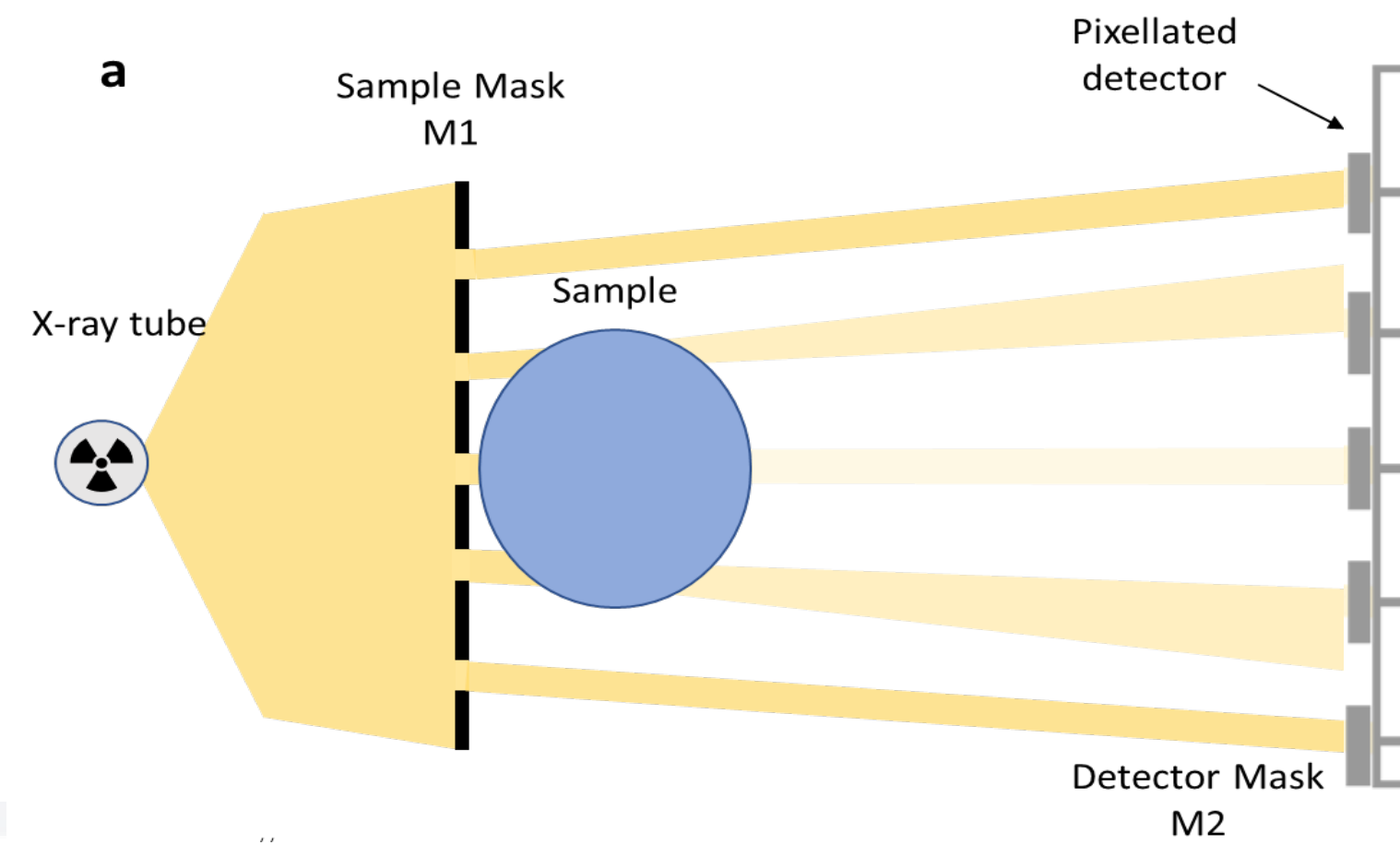
$$\mu_{low} = \frac{\mu}{\rho} \Big|_{low}^{water} \rho^{water} + \frac{\mu}{\rho} \Big|_{low}^{calcium} \rho^{calcium} + \frac{\mu}{\rho} \Big|_{low}^{iodine} \rho^{iodine}$$

$$\mu_{high} = \frac{\mu}{\rho} \Big|_{high}^{water} \rho^{water} + \frac{\mu}{\rho} \Big|_{high}^{calcium} \rho^{calcium} + \frac{\mu}{\rho} \Big|_{high}^{iodine} \rho^{iodine}$$

$$\delta = \frac{\delta}{\rho} \Big|_{water} \rho^{water} + \frac{\delta}{\rho} \Big|_{calcium} \rho^{calcium} + \frac{\delta}{\rho} \Big|_{iodine} \rho^{iodine}$$

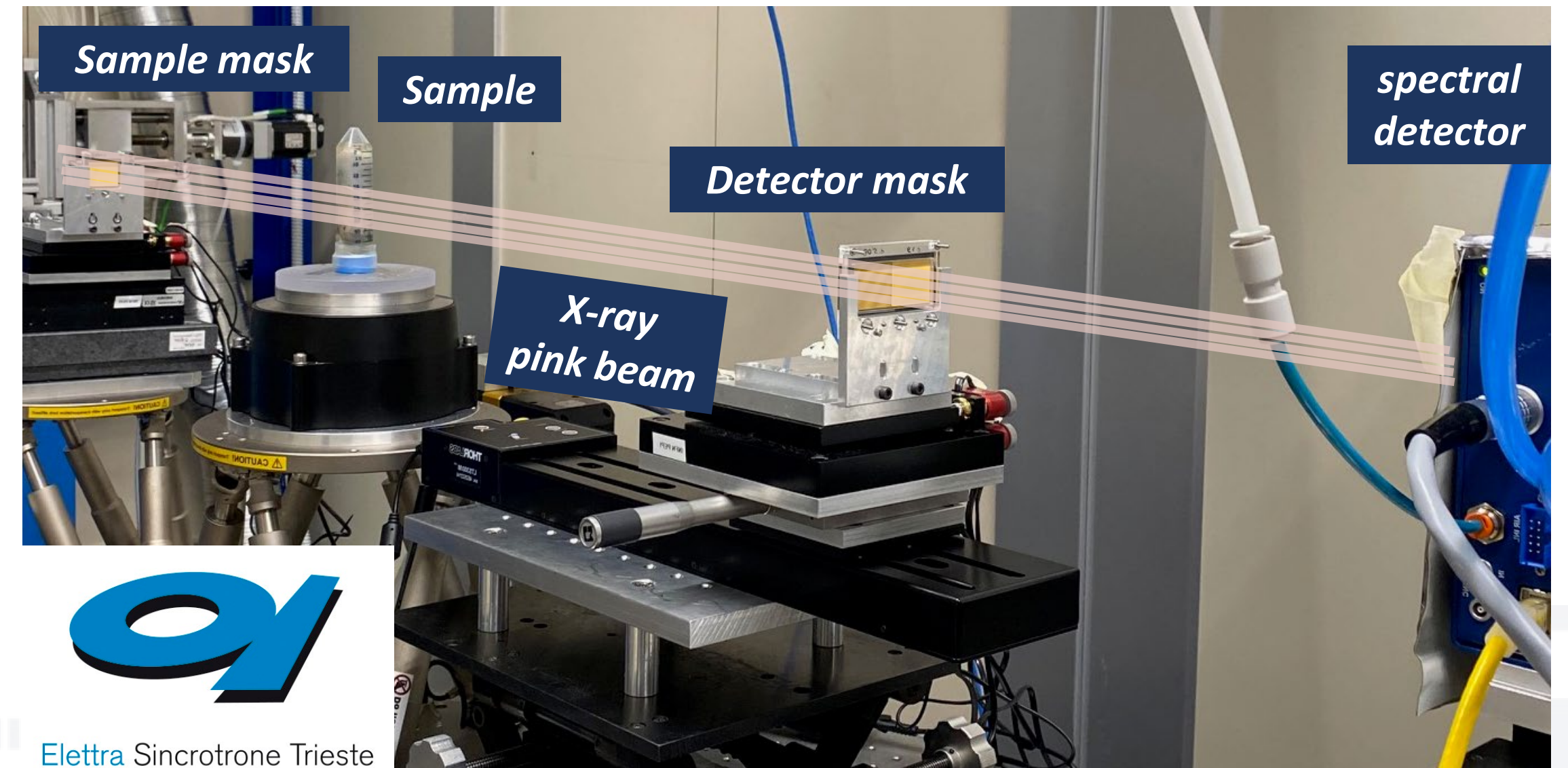
# EDGE-ILLUMINATION SPECTRAL PHASE-CONTRAST CT @ ELETTRA

El technique

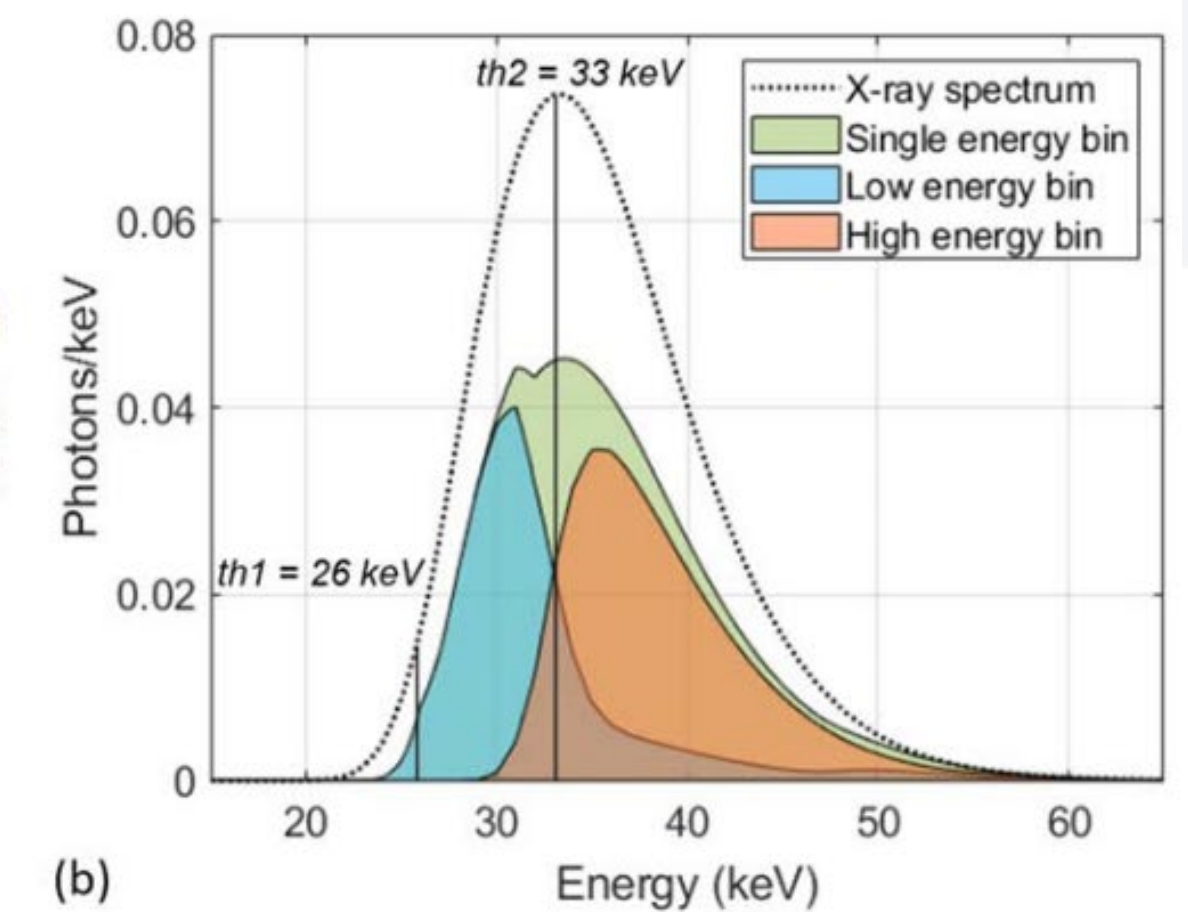
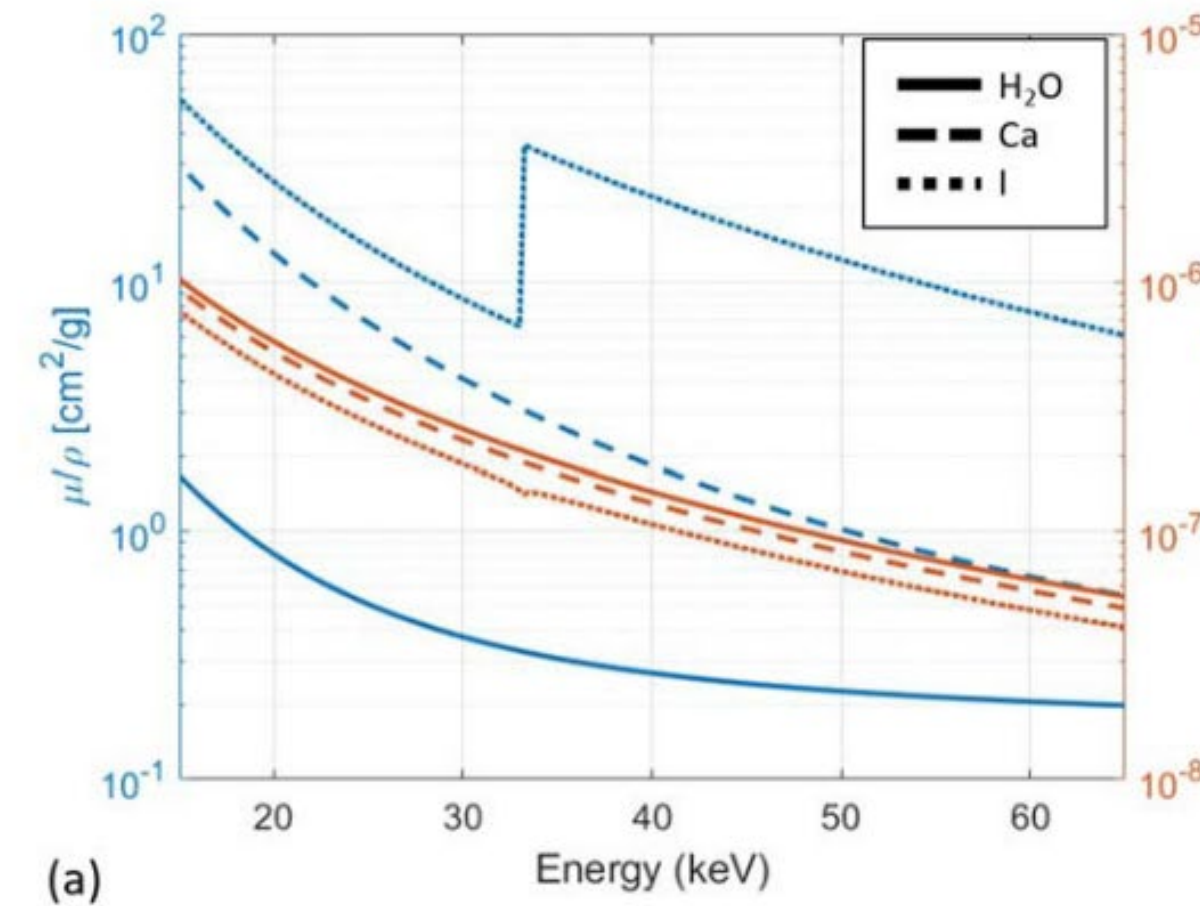


$$IC(x) \rightarrow T \times IC(x - \Delta x) * S$$

Setup

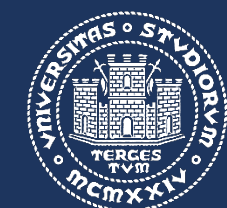


Spectrum

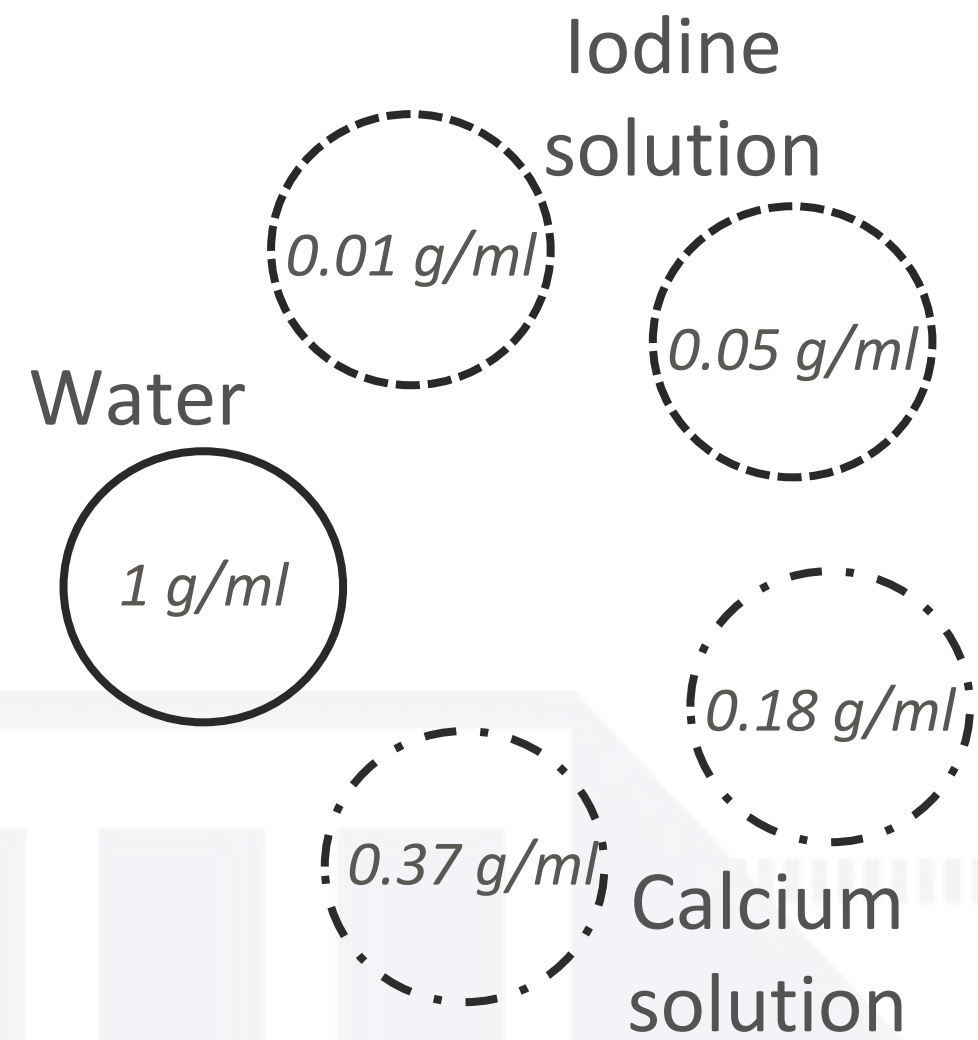


Brombal, Luca, et al. "Edge-illumination spectral phase-contrast tomography." *Physics in Medicine & Biology* 69.7 (2024): 075027.

In collaboration with:



# SPECTRAL

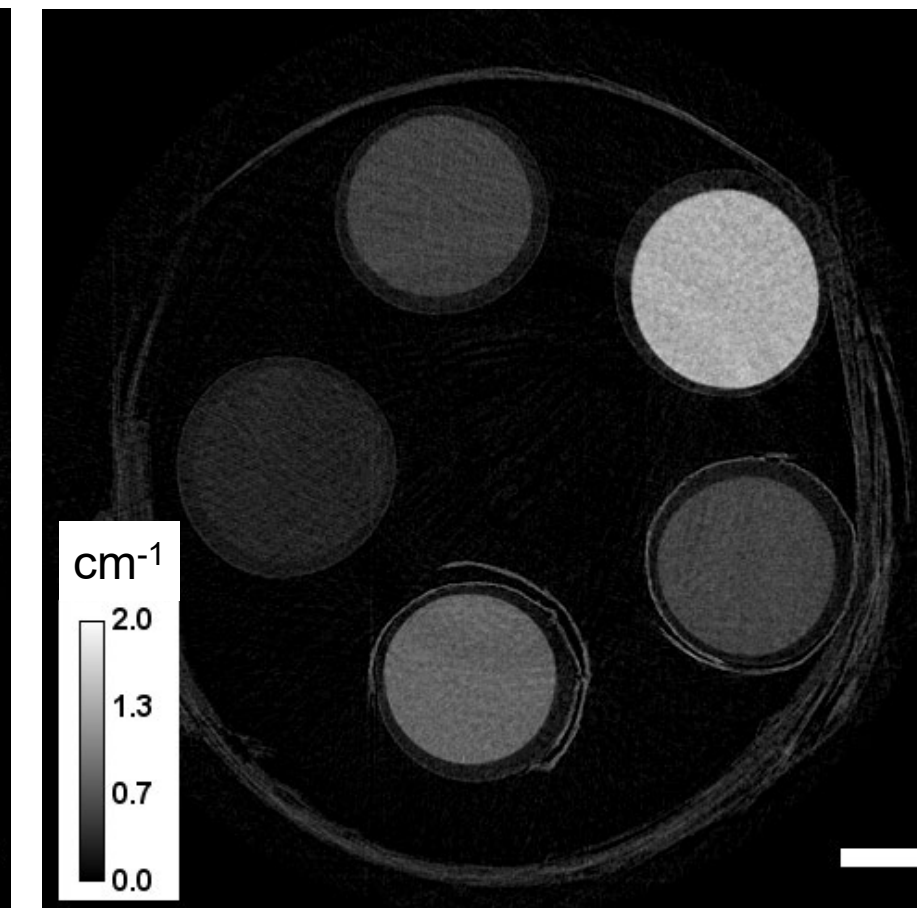
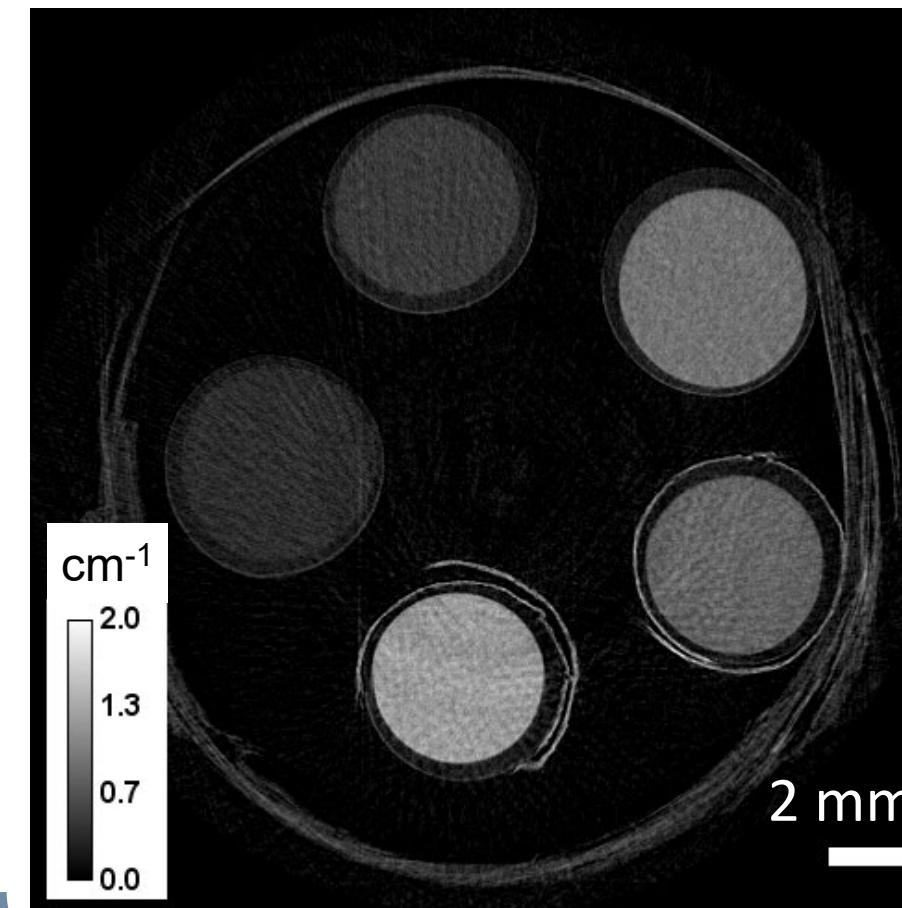


Attenuation  
[26 – 33] keV

Attenuation  
> 33 keV

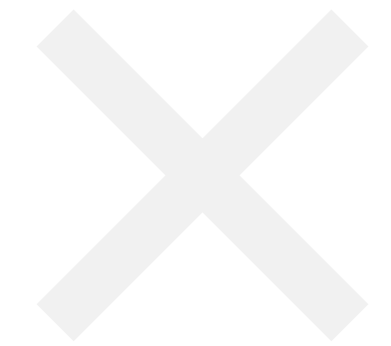
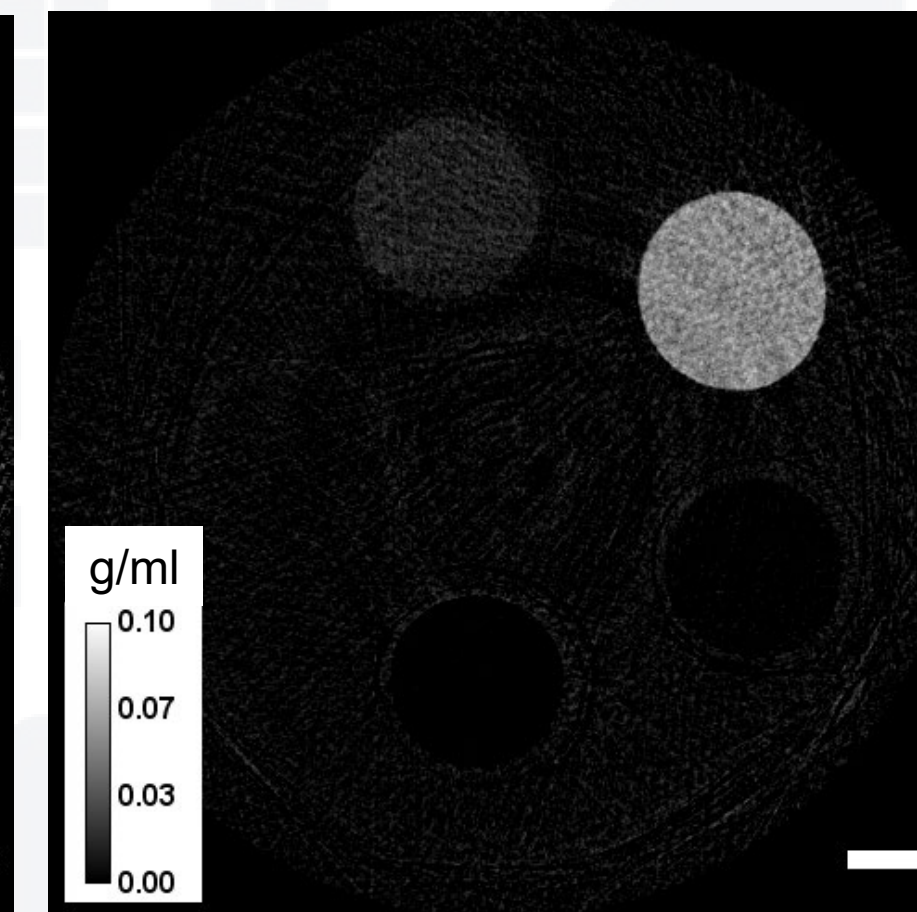
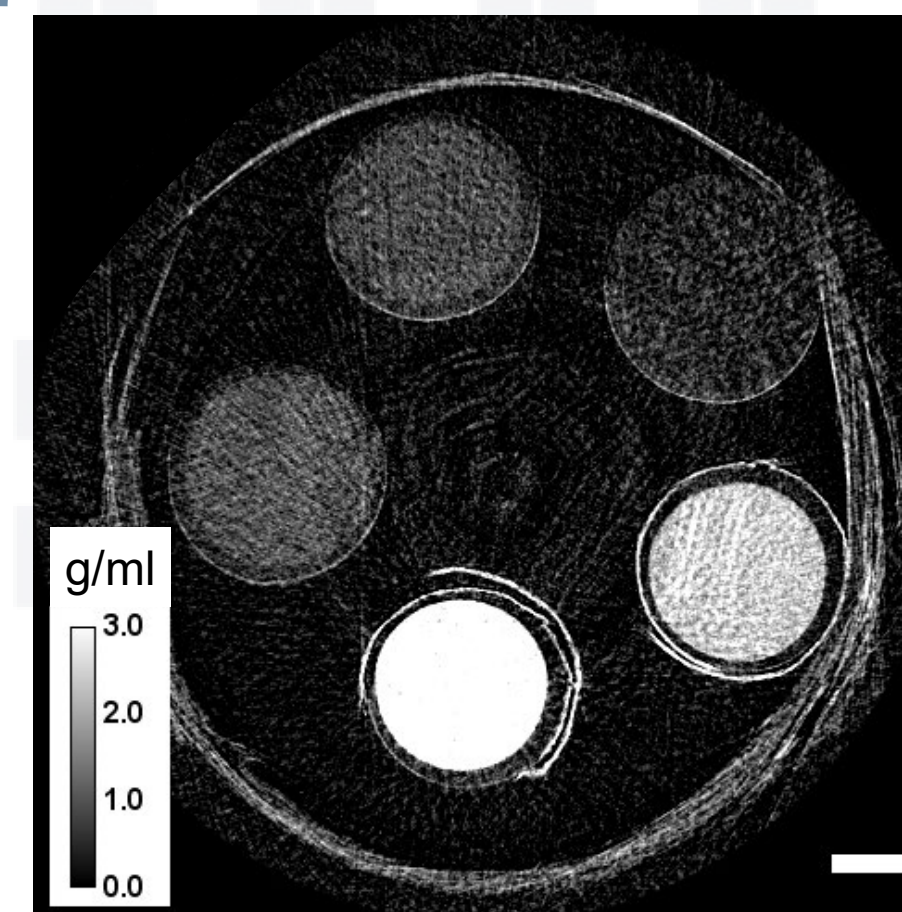
Phase ( $\delta$ )

INPUT



material decomposition

OUTPUT



Water

Iodine

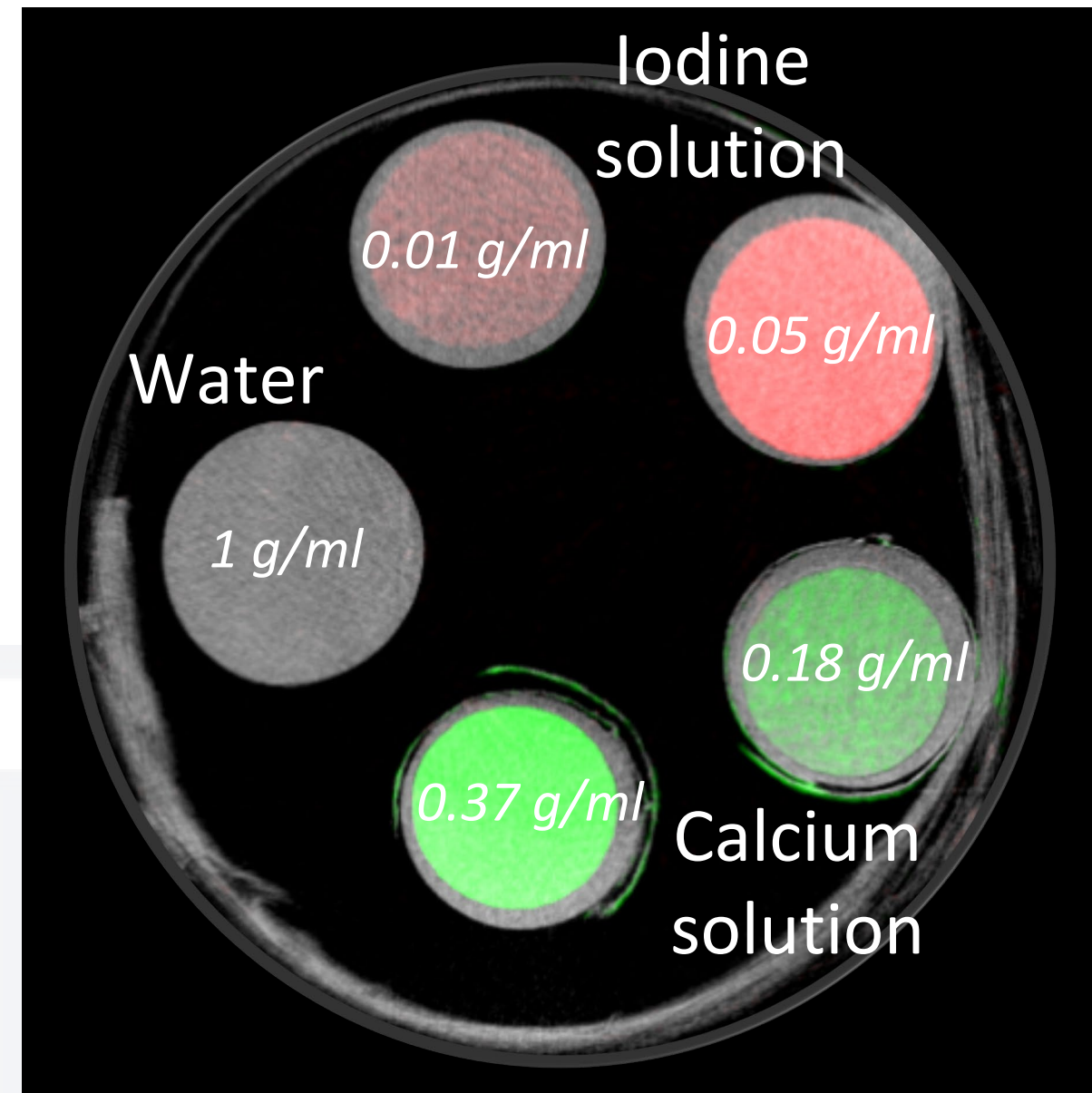
Calcium

- ✓ YES Iodine contrast medium quantification
- × NO calcium quantification
- × High noise in water image
- × Contamination of calcium in water image

Brombal, Luca, et al. "Edge-illumination spectral phase-contrast tomography." *Physics in Medicine & Biology* 69.7 (2024): 075027.

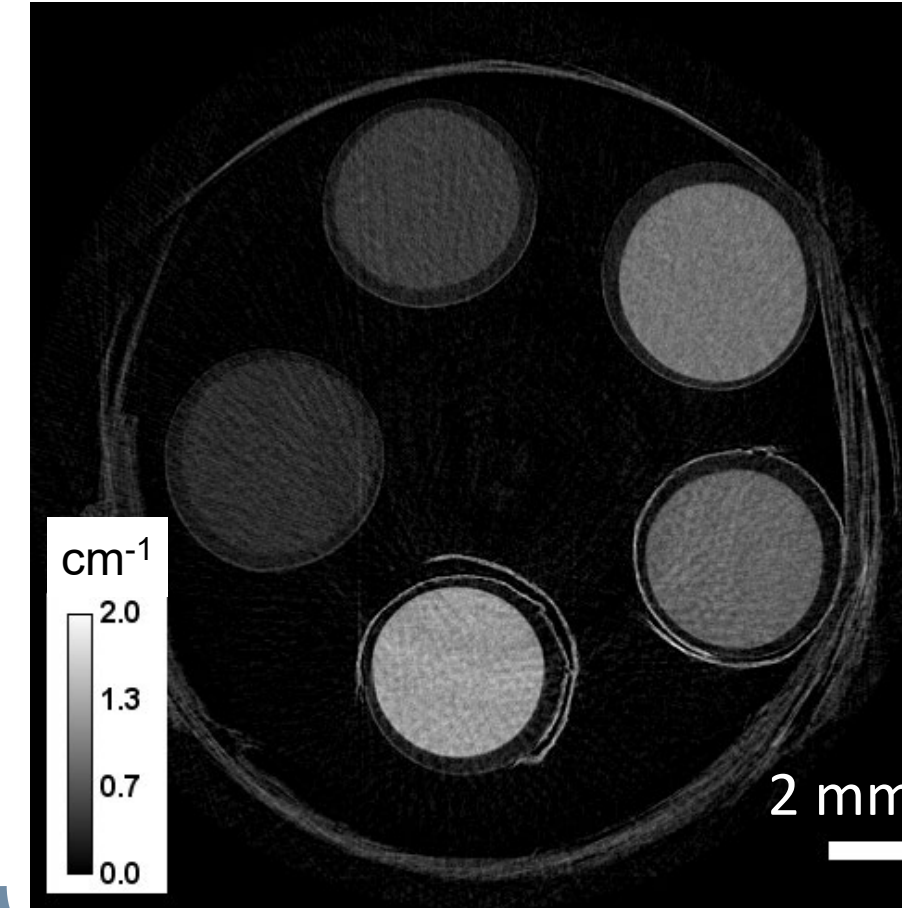


# SPECTRAL + PHASE-CONTRAST

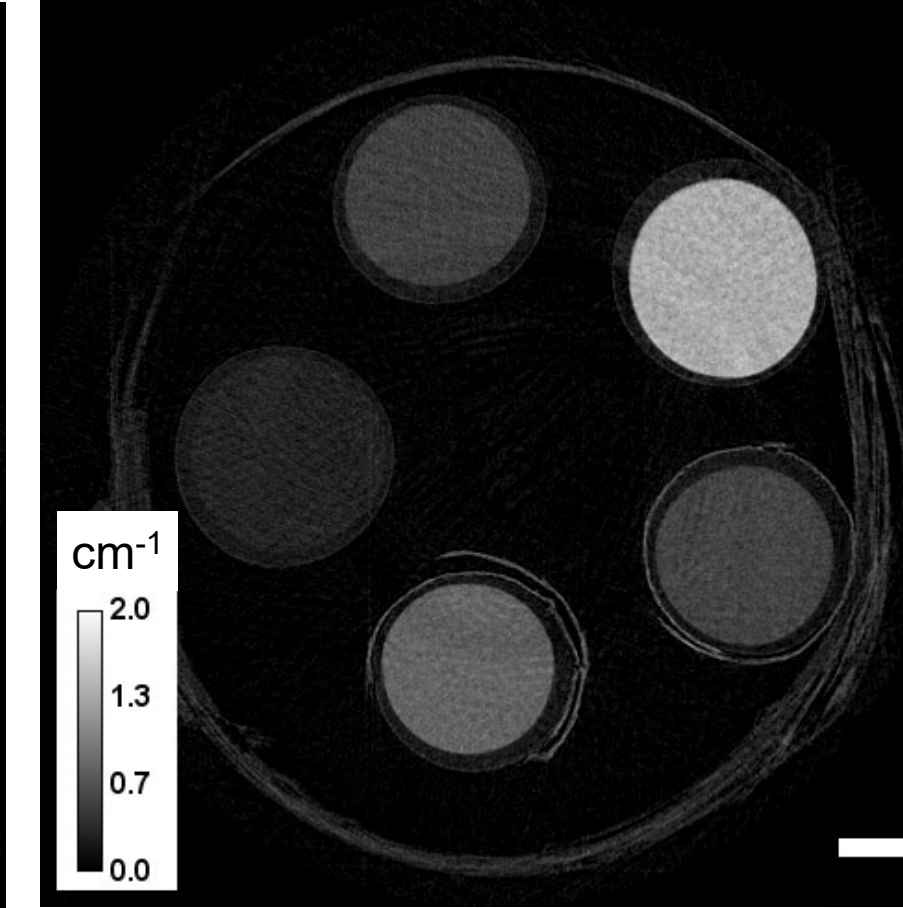


INPUT

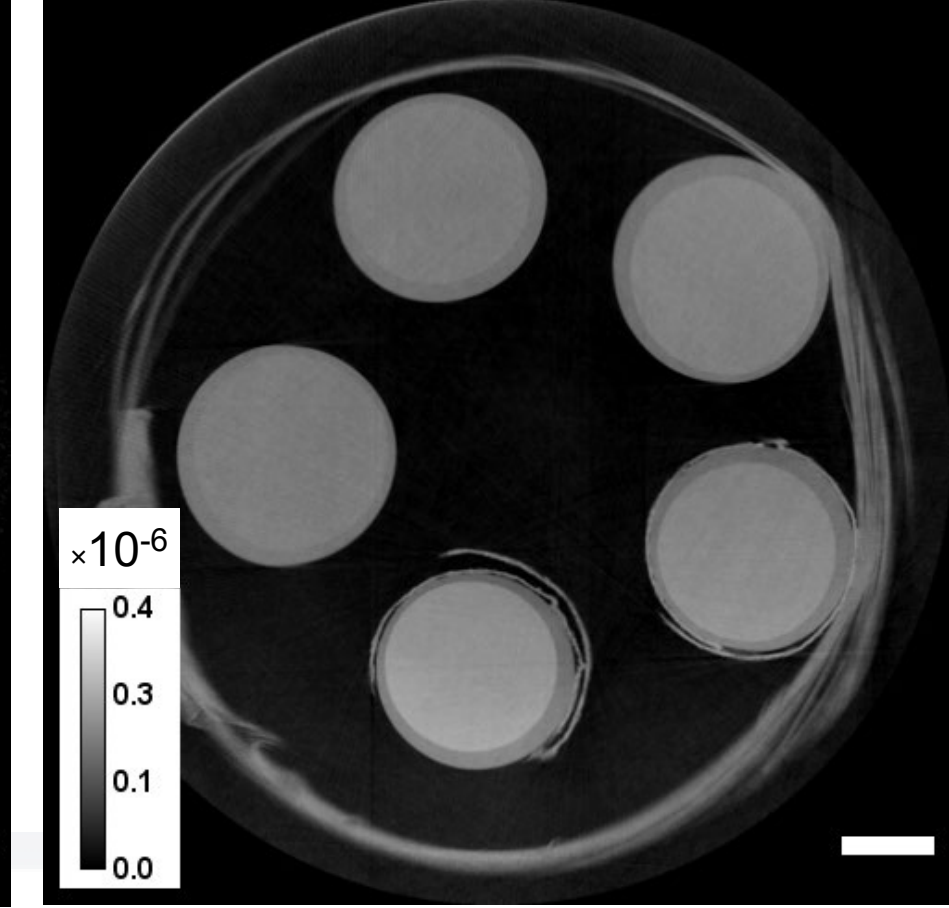
Attenuation  
[26 – 33] keV



Attenuation  
> 33 keV

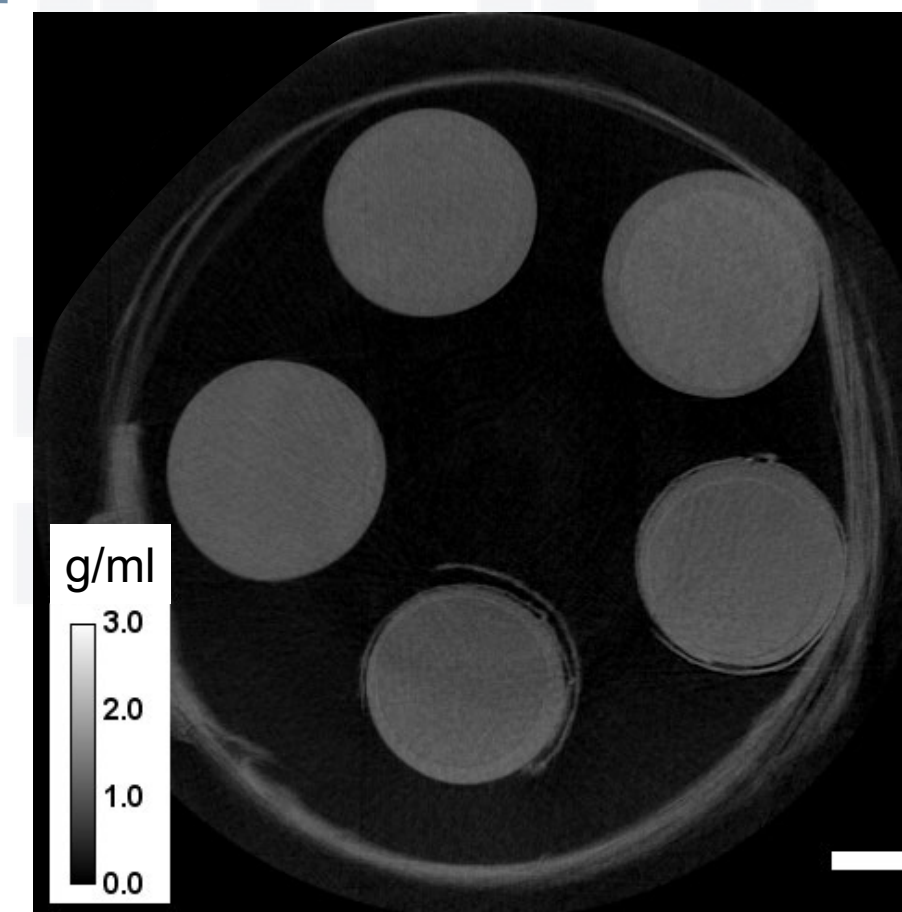


Phase ( $\delta$ )

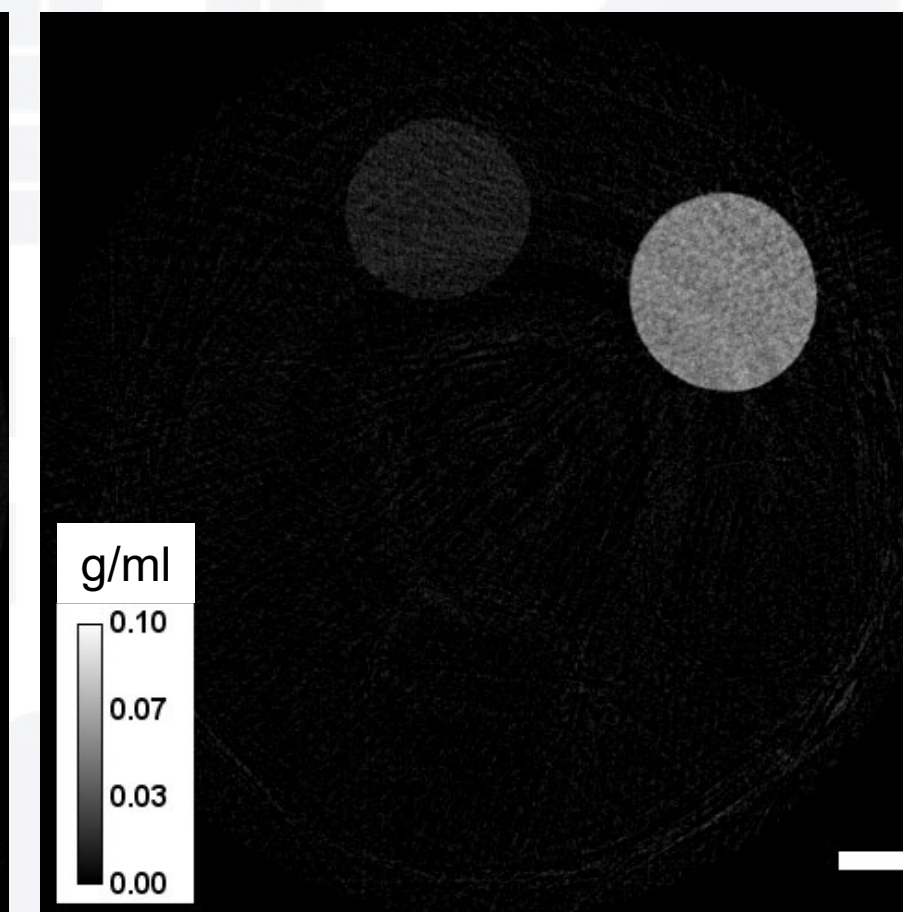


material decomposition

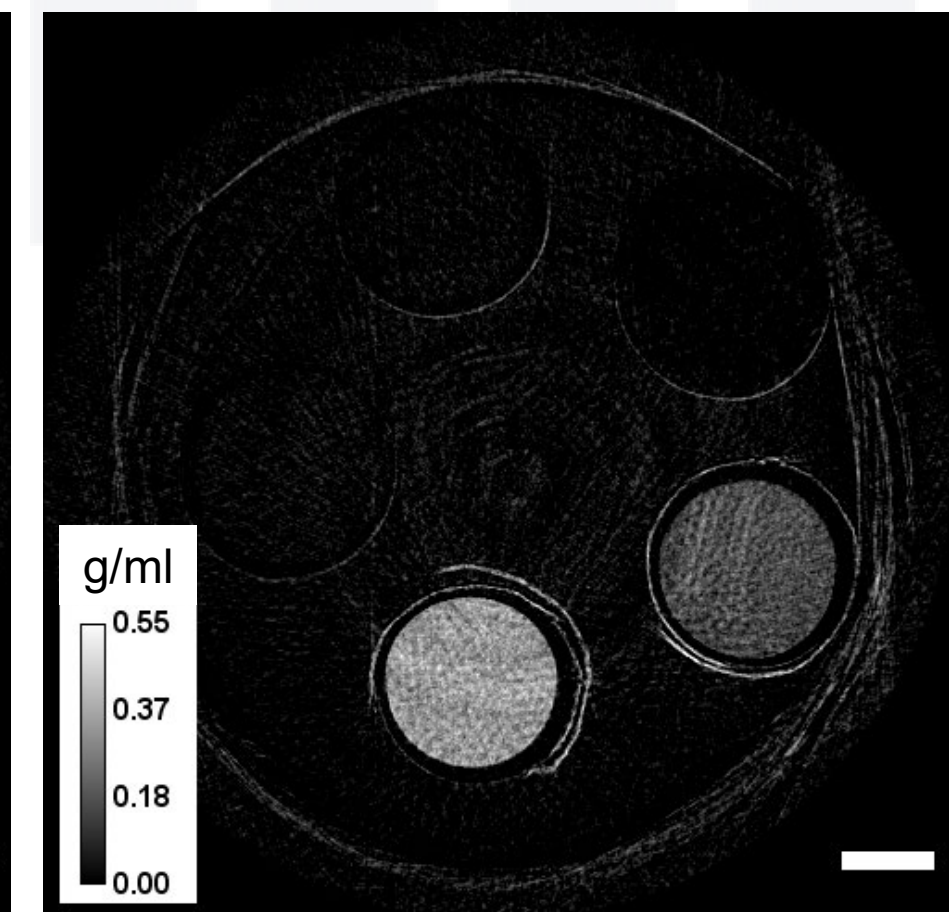
OUTPUT



Water



Iodine



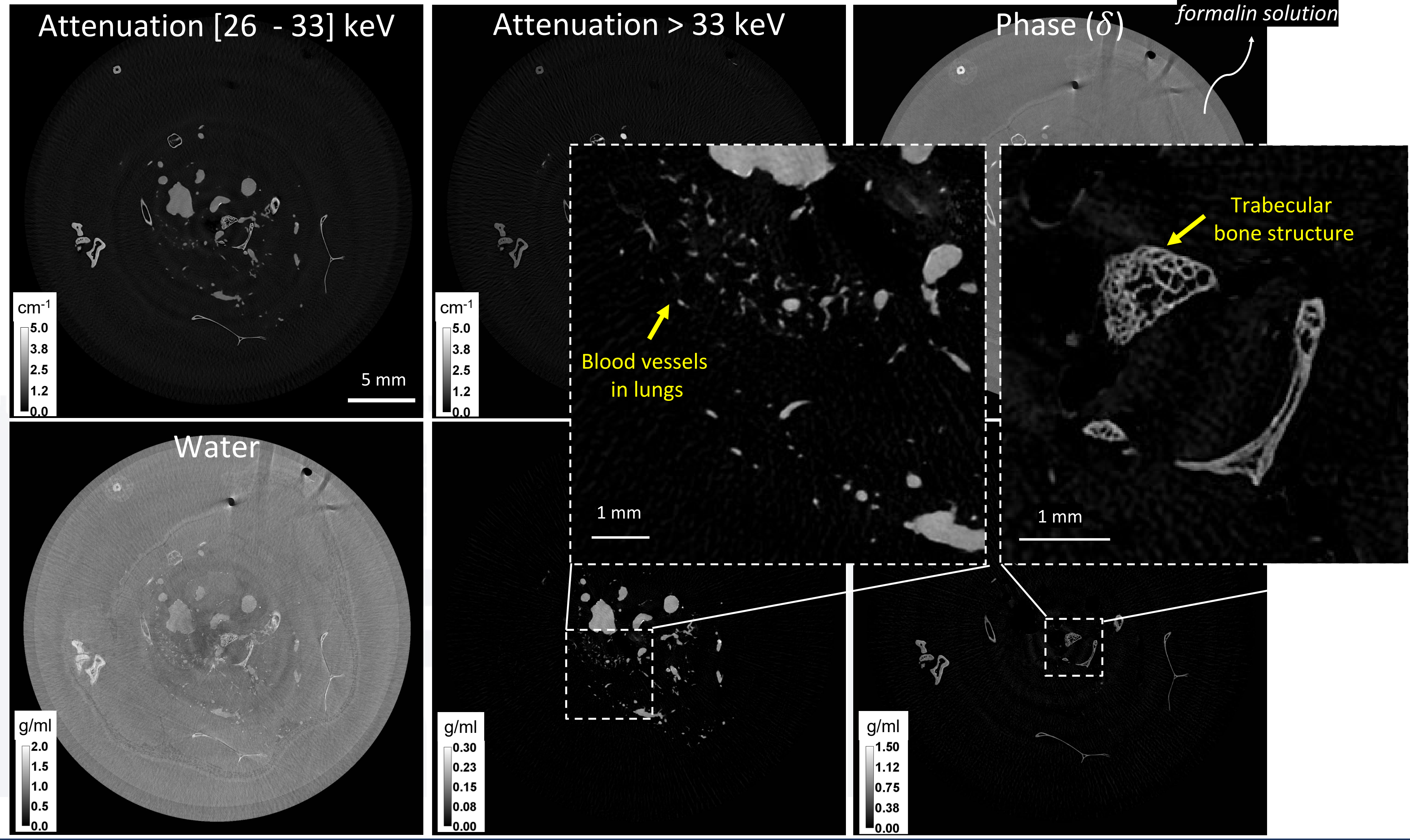
Calcium

- ✓ YES Iodine contrast medium quantification
- ✓ YES Calcium quantification
- ✓ Low noise in decomposed images
- ✓ NO contamination

Brombal, Luca, et al. "Edge-illumination spectral phase-contrast tomography." *Physics in Medicine & Biology* 69.7 (2024): 075027.

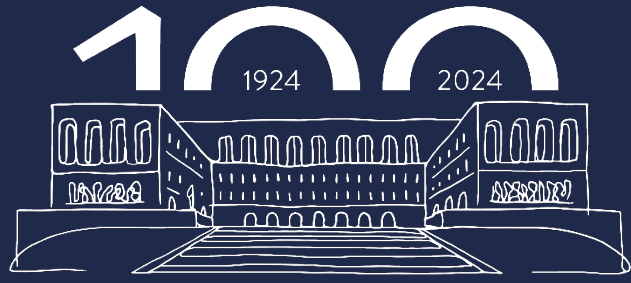
# SPECTRAL PHASE-CONTRAST: EX-VIVO MURINE MODEL

- Murine model (ex-vivo) in formalin
- Perfused (ex-vivo) with iodine-based contrast agent  $\mu$ Angiofil<sup>®</sup>
- In-plane pixel size 20  $\mu\text{m}$



In collaboration with:  
**HELMHOLTZ**  
**MUNICH**

Brombal, Luca, et al. *Phys. Med. Bio.* 69.7 (2024): 075027.



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...TAKE HOME...

- Detector-based spectral imaging already in clinics and in micro-CT laboratories
- Phase-contrast is widely used in pre-clinical and non-clinical studies. It is moving its first step into clinics
- The combination of the two techniques will become an option in the future and research is starting now



Finanziato  
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NextGenerationEU

THANK YOU!