

Pinhole X-ray Fluorescence Imaging of Gadolinium Nanoparticles :A Preliminary Monte Carlo Study

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Contents

- Introduction
- Material and Methods
- Results
- Conclusions
- Future Work
- Q&A

Introduction

- Metal nanoparticles in medical applications

1. Metal Enhanced Radiation Therapy



- Dose enhancement due to photoelectrons/Auger electrons of High Z material
 - 140kVp source
 - a. 3% gold in tissue → 560% dose enhancement
 - b. 0.7% gold in tissue → 211% dose enhancement
- (Cho S.H., 2005, PMB)

2. Molecular Imaging

Tumor targeting metal nanoparticles as a molecular probe

- Molecular size
- Functional imaging


Introduction

- X-ray fluorescence imaging
 1. *in vivo* imaging of metal nanoparticles distributed within tumor and other critical organ  **Location**
 2. *in vivo* quantification of the amount of metal nanoparticles  **Concentration**

Actual accumulated concentration of nanoparticles

- The expected concentration of gold accumulated in tumors is on **the order of 0.001%** (*J.Hainfeld et al., BJR, 2011*)
- In case of Cisplatin chemotherapy, **concentrations of 0.0005% ~ 0.004% have been measured** (*R. Jonson et al., Acta Oncol, 1991*)

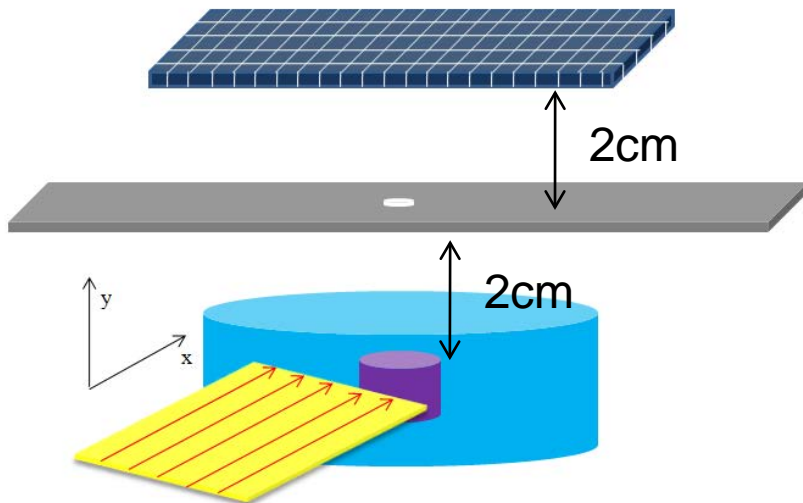
Introduction

- Purpose
 1. Imaging **relatively low concentrations** of **gadolinium nanoparticles** using (1)**x-ray fluorescence**, (2)**pinhole** imaging system and (3)**monochromatic** x ray source
 2. Comparing two energy (1)**50.5keV**, (2)**55keV** of **photon source**
- **Gadolinium(Z=64)**
 - Most widely used metal nanoparticles in medical applications
 - MR contrast
 - Dose enhancer in radiation therapy
 - Kedge energy: 50.239keV
 - Characteristic X-rays: $K_{\alpha 1}$ (42.996keV), $K_{\alpha 2}$ (42.309keV),...
 - Kedge $E < 60\text{keV}$ (포항광가속기에서 조사 가능한 최대 mono Energy)
 - Toxicity  concentration of Gd injected should be low

Material and Methods

- **Pinhole** imaging system
 1. Magnification, high spatial resolution
 2. 2D radiography with 2D detector array
 3. Benchmarking an image modality pinhole SPECT(Single Photon Emission Computed Tomography)

- Monte Carlo simulation geometry



Dexel: $0.25 \times 0.25 \text{cm}^2$
 Detector array: $4.5 \times 4.5 \text{cm}^2$

Pinhole: 0.4cm diameter

Source: 1mm height rectangular shaped x-ray beam

Water phantom: 5cm diameter
 Gd column: 1cm diameter

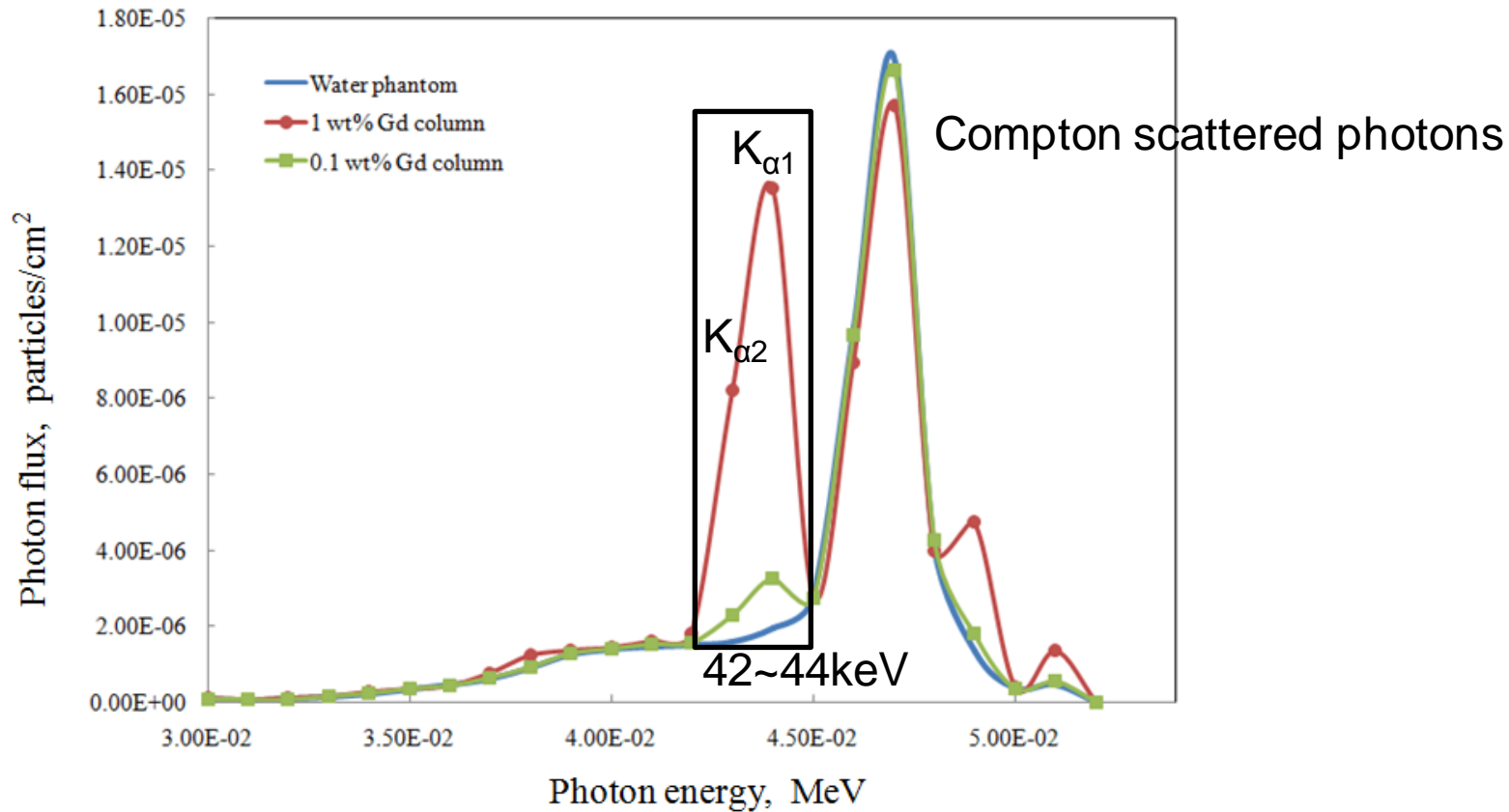
Material and Methods

- Monte Carlo simulation
 - MCNP6
 - Photon only, phys:default, E cutoff: 41.9keV, no variance reduction
 - Flux and energy bin tally of each detector pixel
 - Nps 2.0E10
 - Uncertainty < 5%

- Simulation conducted for
 - Water phantom
 - Single column inserted water phantom: 0.001%~0.1% by weight of Gd
 - Three columns inserted water phantom: 0.1-0.25-0.5,
0.05-0.075-0.1 % by weight of Gd

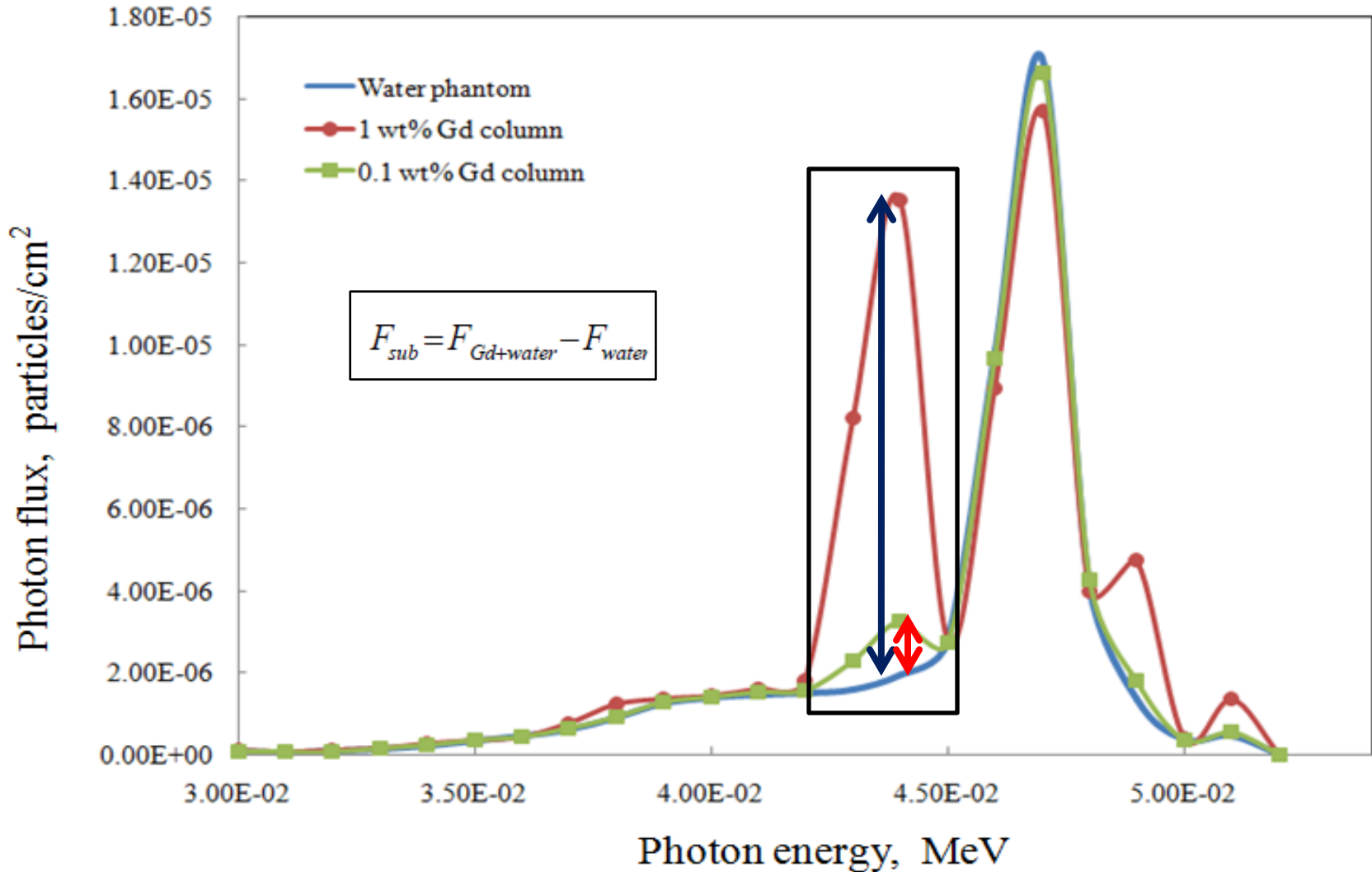
Material and Methods

- Data acquisition
 - 50.5keV source
 - energy spectrum of **central pixel** with and without Gd column



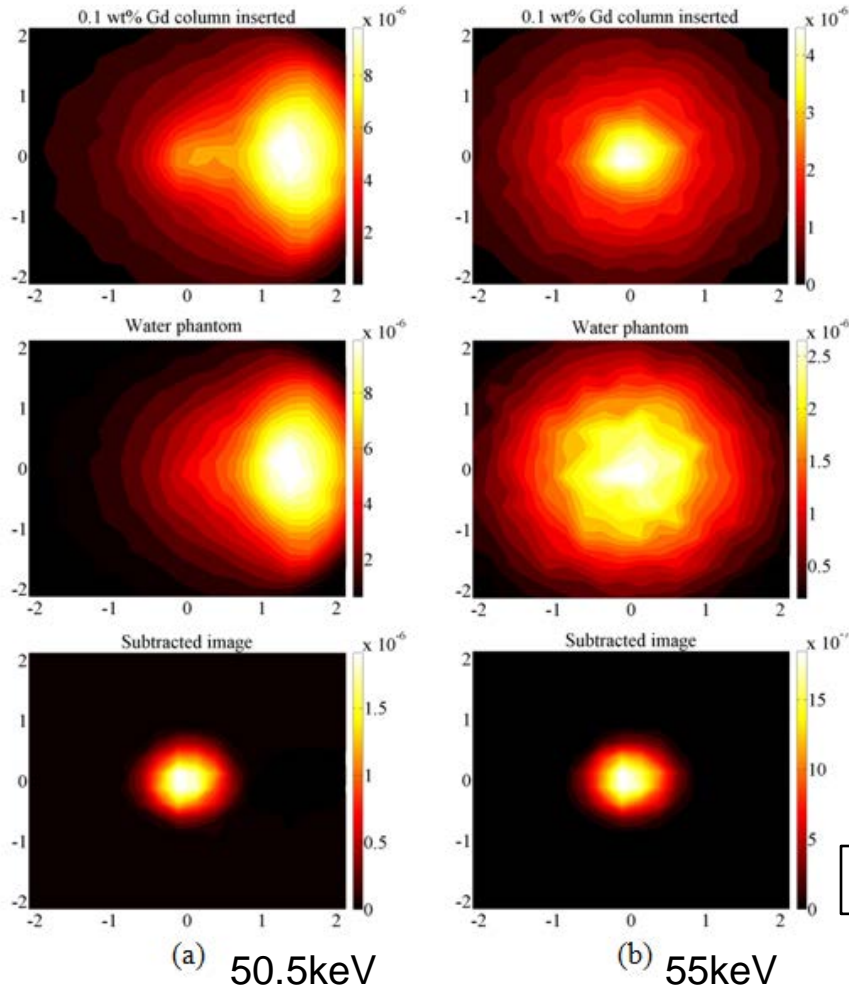
Material and Methods

- Data acquisition



Material and Methods

- Data acquisition



(a) 50.5keV
The effect of Compton scattered photons is bigger than 55keV

$$E' = \frac{E_{\gamma}}{1 + \frac{E_{\gamma}}{m_0c^2}(1 - \cos \theta)}$$

If $E'=43\text{keV}$, $E_{\gamma}=50.5\text{keV}$, then $\Theta = 140^{\circ}$
If $E'=43\text{keV}$, $E_{\gamma}=55\text{keV}$, then Θ does not exist

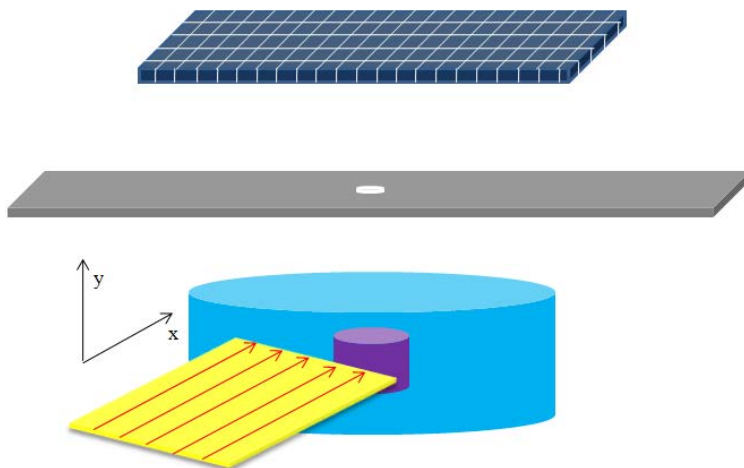
0.1 wt% Gd

Material and Methods

- Image processing

1. Attenuation correction, k_a

- x direction: the primary beam through the phantom
- y direction: Gd fluorescence *en route* to the detector



$$k_a = \frac{1}{e^{-\mu_{w,E} \cdot x} \cdot e^{-\mu_{w,K} \cdot y}}$$

$\mu_{w,E}$: Linear attenuation coefficient of water for photons of energy E

$\mu_{w,K}$: Linear attenuation coefficient of water for photons of K characteristic x-rays

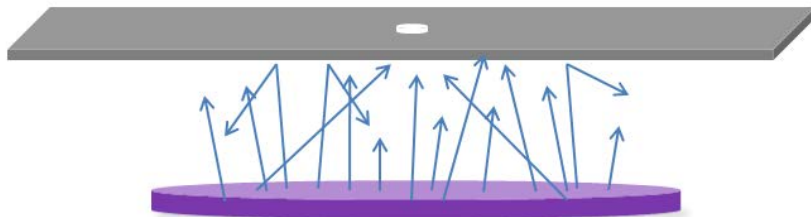
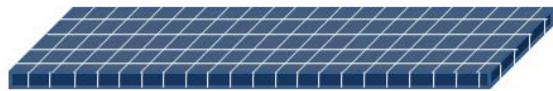
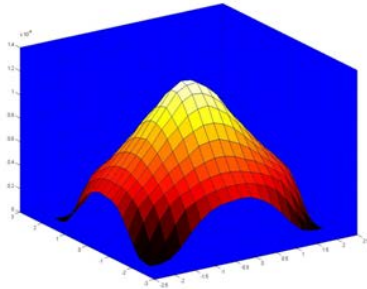
x: the path length of photons before photoelectric interaction

y: the path length of fluorescence photons after the interactions

Material and Methods

- Image processing

2. Inverse correction, k_r



K characteristic like 43keV
isotropic photon emitting
disc source

- Inverse r square law
- Point spread function

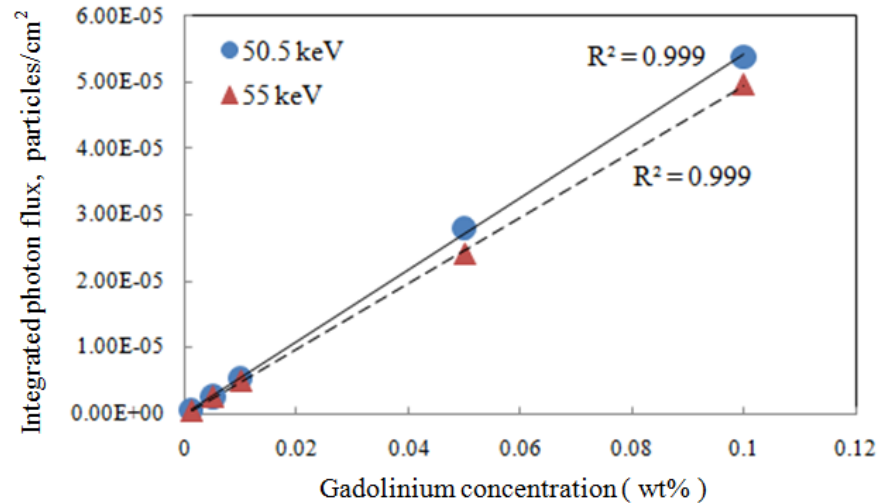
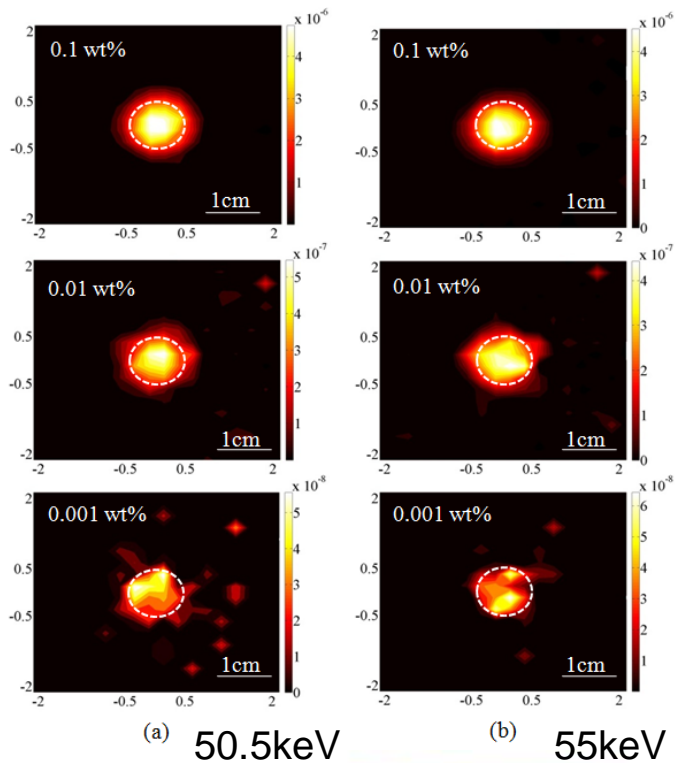
➡ could inversely get a correction factor from the left Monte Carlo simulation

$$F = F_{sub} \cdot k_a \cdot k_r$$

Pixel value of image

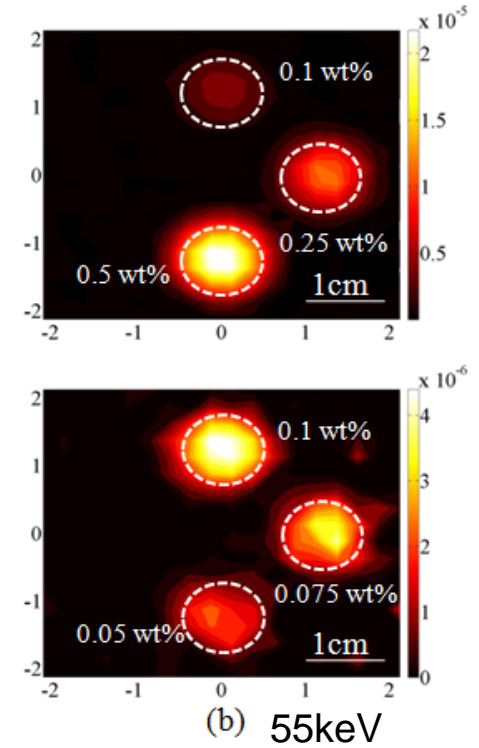
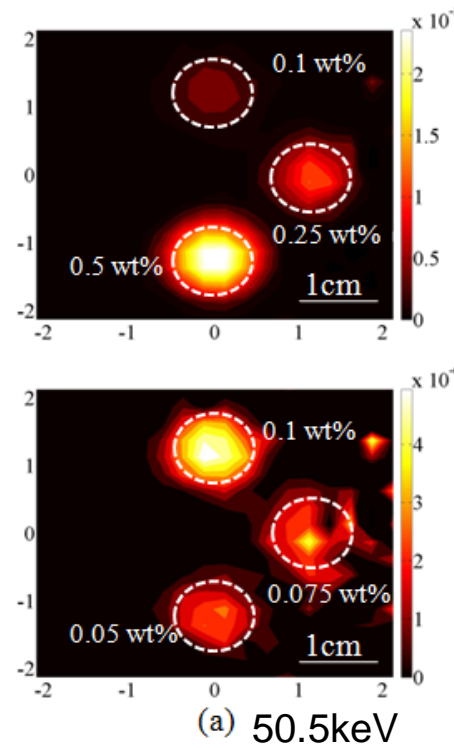
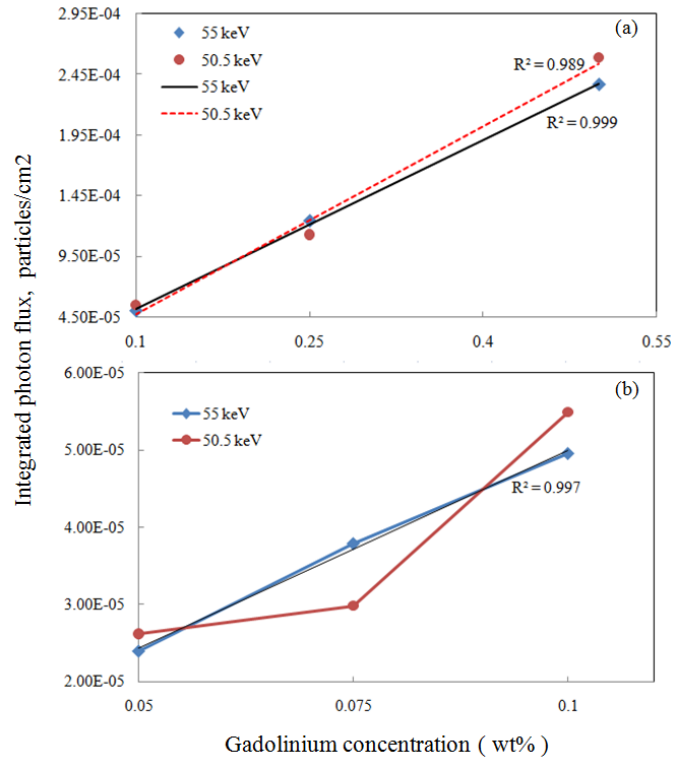
Results

- Relation between concentration and F value
 - Integrated photon flux
 - F values of the pixels expected to detect Gd nanoparticles could be integrated, since we know the locations of Gd columns
- Single column, 0.001~0.1 wt% Gd



Results

- Three columns



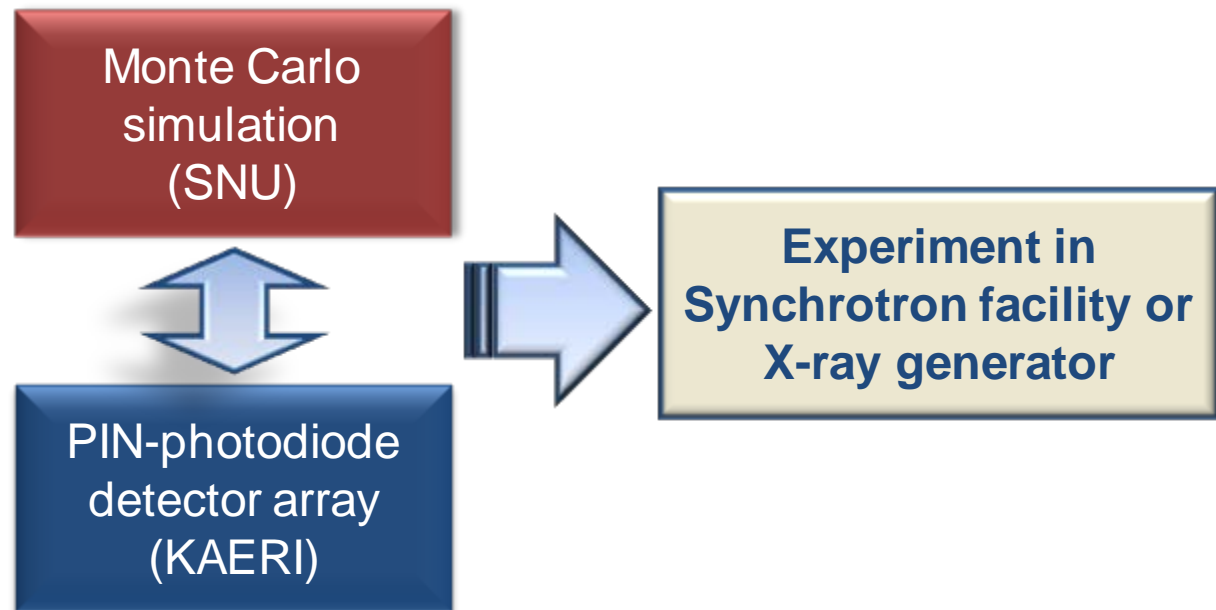
Conclusion

- Using Monte Carlo simulations, **the feasibility of imaging low concentrations of Gd nanoparticles** with x-ray fluorescence using monochromatic synchrotron x-rays of two different energies was shown
- The photon beam of 55keV showed better images and linear relationship between the three different concentrations and locations Gd columns
- Due to the region of Compton scattering noise, images of Gd columns irradiated by 50.5keV failed to make a linear relationship

Future Work

- More simulations are needed for Gd columns with lower concentrations
- 2D radiography(current study) → 2D reconstruction image
- Imaging metal nanoparticles during Microbeam Radiation Therapy(MRT)
- Benchmarking multi-pinhole SPECT
- Simulations for magnified images
 - High spatial resolution

- Experiment





Q&A

Thank you