

Theranostics and Personalized Therapies in Nuclear Medicine

Carlos Uribe, PhD, MCCPM

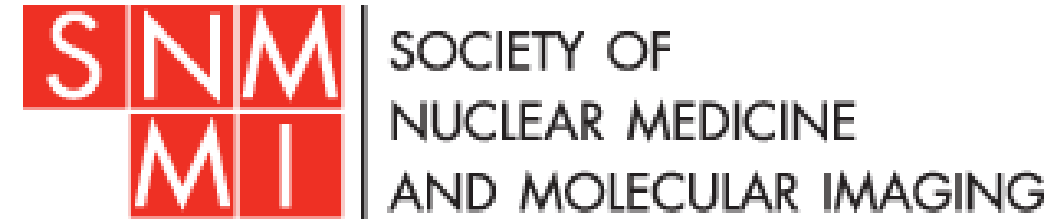
December 13, 2021



What is Nuclear Medicine?

What is Nuclear Medicine?

- Medical Specialty used to
 - Diagnose
 - Treat
- Allows to gather medical information that may otherwise:
 - Require Surgery
 - Require more expensive and invasive diagnostic exams
- It can identify abnormalities very early in the progression of the disease (even before they are apparent with other diagnostic tests)
 - Earlier treatment -> higher chances of a better prognosis



Radioactivity in Medicine

Generate images
of an organ

- Radioactive tracers are injected into the patient
 - Planar (2D)
 - Tomographic (3D)

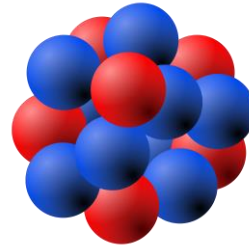
Therapy

- Radioactive tracers are injected into the patient to deliver energy and hopefully kill cancer cells (tumors).

Why is it called “Nuclear”?

A medicine attached to a small quantity of radioactive material

Pharmaceutical + **Radioisotope** = **Radiopharmaceutical**



Radiopharmaceuticals

- Link between physics (radio) and biochemistry (tracer)
- Is a substance that is labeled with a radioisotope, that decays allowing us to determine its location
- Inside the patient, it has a spatial/temporal distribution that provides useful information (e.g. For diagnosis)
 - The distribution or uptake depends on the biochemical properties of the tracer, not the fact that is radioactive
 - Necrotic tissue does not have tracer uptake
 - Uptake shows tissue function
- Nuclear medicine falls below the “functional imaging” category. It does not show anatomy or structure.

Tracers and Radioisotopes

PET Isotopes (Positron Emitters)

Half-life	Rmean (mm)	Isotope
110 min	0.6	^{18}F
68 min	3.5	^{68}Ga
20.4 min	1.2	^{11}C
78.4 h	1.3	^{89}Zr

Therapy Isotopes (Beta emitters)

Half-life	Rmean (mm)	Gamma emissions [keV]	Isotope
6.7 days	0.5	113 (6.2) 208 (10.3)	^{177}Lu

Small molecules
(Deoxyglucose)

Peptides and derivatives
(DCFPyL)
(PSMA-617)

Monoclonal antibodies
(Rituximab)

Role of Physics in Nuclear Medicine

- Underlying physics is not changing
- But the technology is changing
 - Production of radioisotopes (with the radiochemist)
 - New detectors, new configurations
- Methods for accurately quantifying concentrations of radionuclides within the patient
- Improvement of the models used for radiation dosimetry
- Physics plays a role in providing high-quality, cost-effective, quantitative, reliable, and biologically safe assays in Nuclear Medicine.

Nuclear Medicine can help in

- Determining if an organ is functioning well, or if there is any disease.
- Establishing if the physiology and/or metabolism has been altered?
- Providing information about treatment efficacy.
- Drug development
- Classification of disease
- Reporting of gene expression
- ... and many more

Emission Detection

Groups of Radiation Detectors

We are interested in detecting gammas emitted by radiopharmaceuticals to generate images

The processes of photon interaction with matter result in production of energetic electrons that transfer their energy to the medium by ionization and excitation

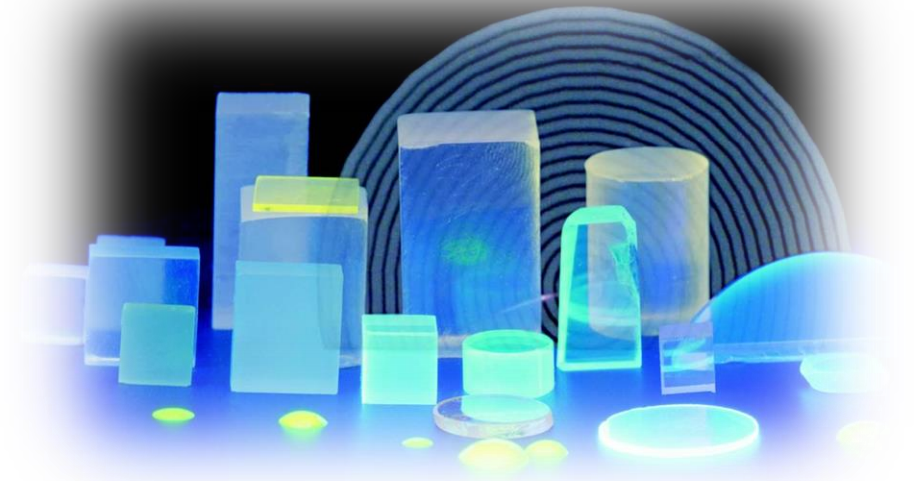
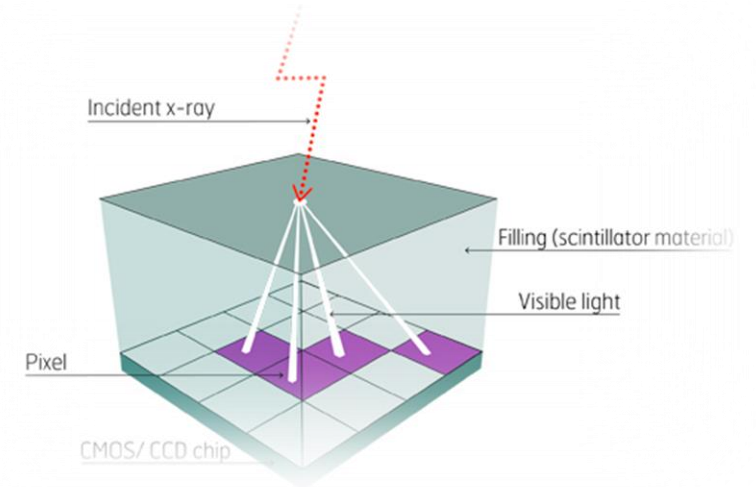
Charge carriers
Gas detectors

Electrons and holes
Semiconductor detectors

Emission of light
Scintillators

Scintillators

- Some materials emit light when they interact with ionizing radiation
- Scintillators can be plastic, organic or inorganic crystals, liquids or gases.



Scintillator Detectors

Inorganic ionic crystals are the most important in Nuclear Medicine

- High density and Z
- Fast response and high light yield
- Large crystals can be grown

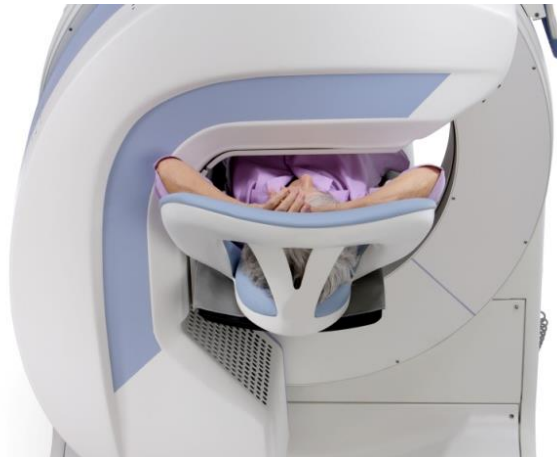
Important property of scintillators

- **Amount of light produced after the interaction with a particle is proportional to the energy deposited by the particle in the scintillator.** In the NM energy range, the amount of light is very small.

Scintillator Detectors

Scintillator	Light Yield (photons/MeV)	Density (g/cm ³)	Hydroscopic?	Decay constant (ns)	Energy resolution (%)	Max emission wavelength (nm)
NaI	41000	3.7	Yes	230	7	410
BGO	9000	7.1	No	300	21	480
LSO	26000	7.4	No	40	12	420
LYSO	26000	7.3	No	50	13	420
GSO	8000	6.7	No	60	14	440
BaF ₂	1400	4.9	No	0.8	10	225

The Gamma Camera

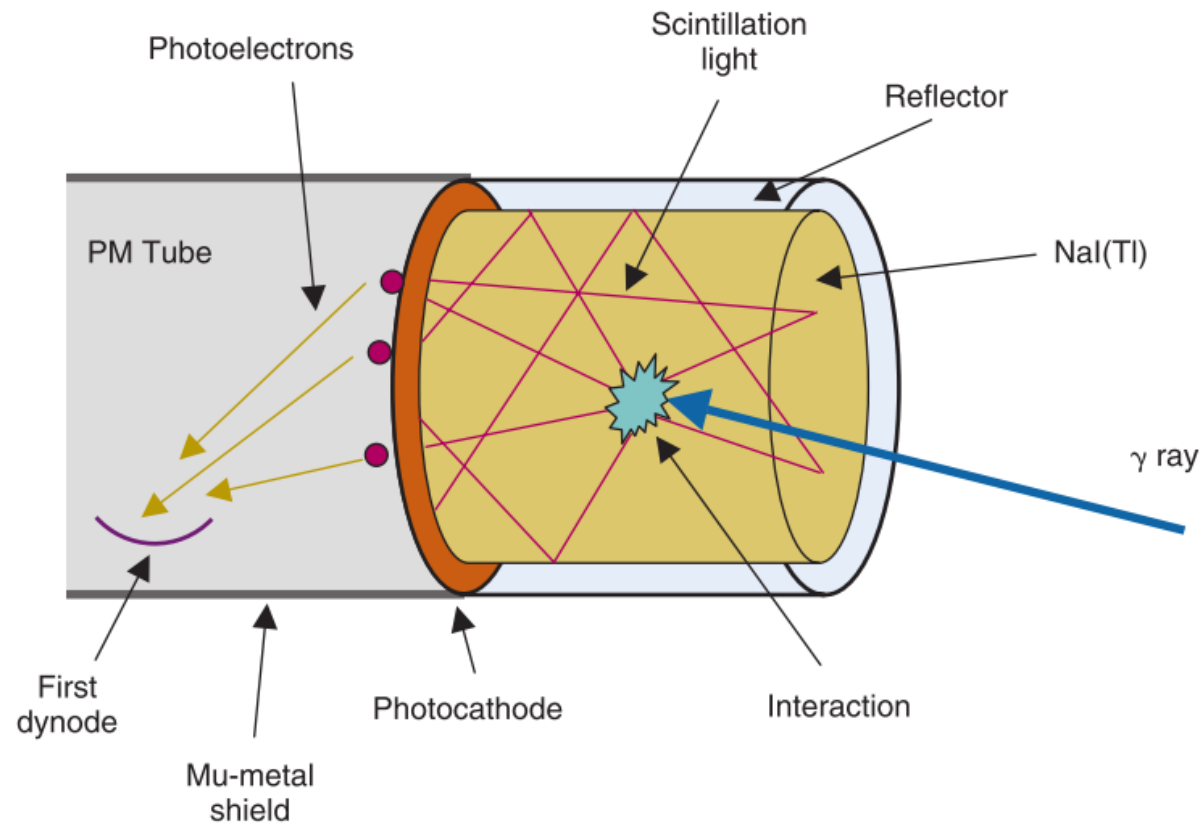


Single Photon Emission Computed Tomography
(SPECT)

The Gamma Camera

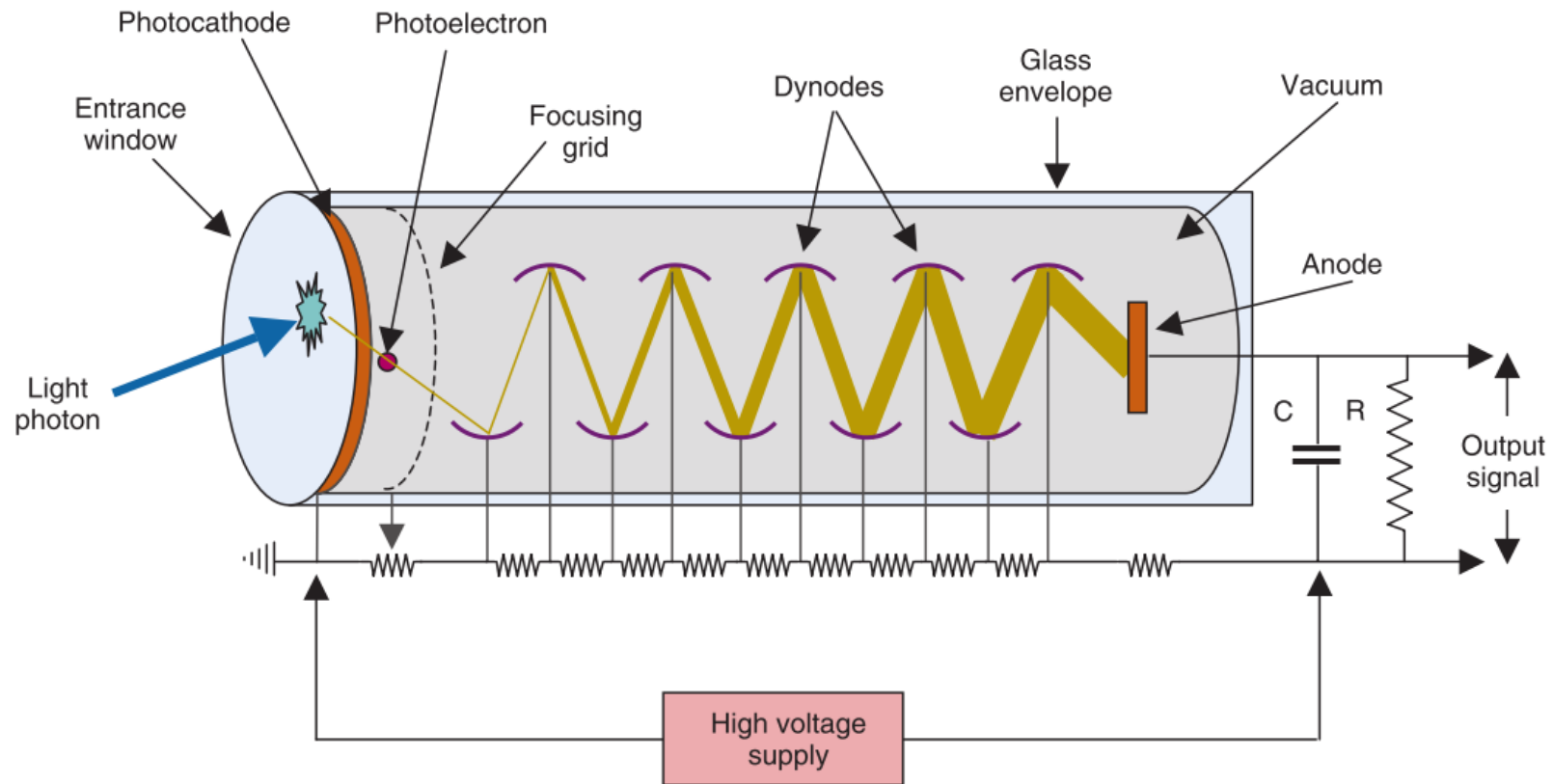
- Detection of gammas from single photon emitting radioisotope
 - Energy of gammas typically within 70-400 keV. (Tc-99m -> 140 keV)
 - The energy depends on the isotope
- Statistics → Record as many photons as possible
- Spatial resolution → localization of photons during detection
- Energy resolution → reject events that do not correspond to the source.
- Isotope identification
- The direction from which the photons reach the detector matters

The Gamma Camera



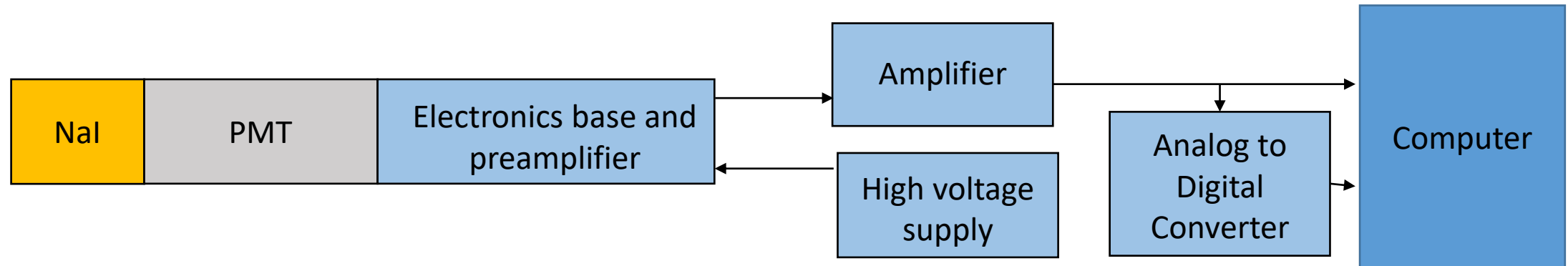
Cherry, Physics in Nuclear Medicine

The Gamma Camera

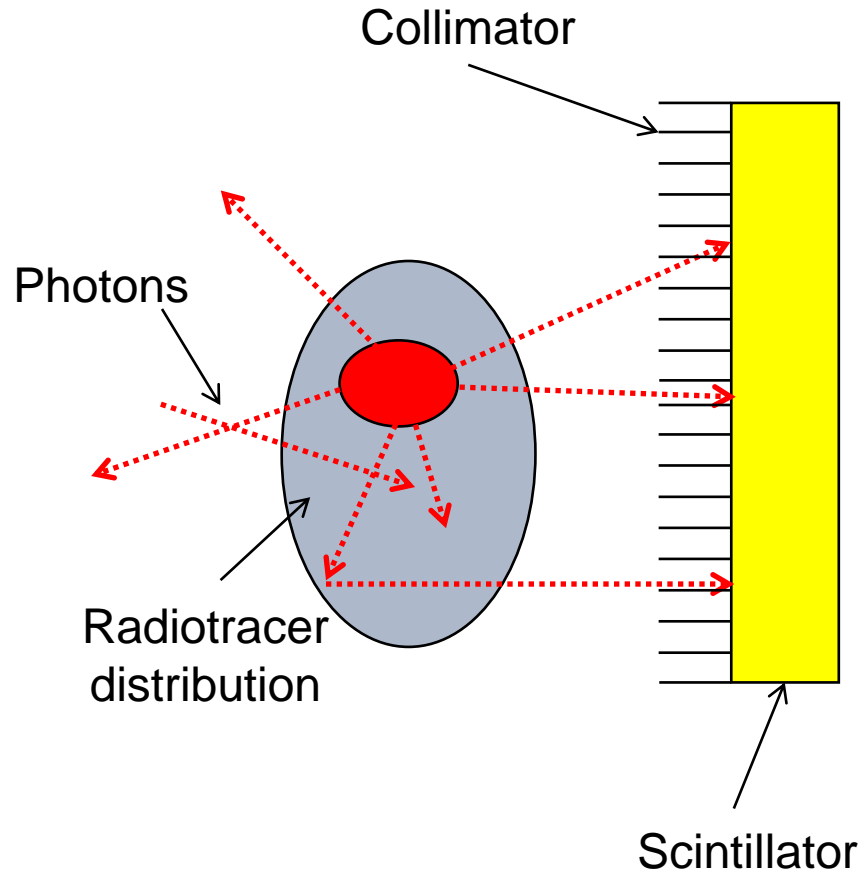


Cherry, Physics in Nuclear Medicine

The Gamma Camera



Absorptive Collimation



Gamma rays cannot be focused with lenses as it is done in cameras

Collimators allow only those photons travelling along certain directions to reach the detector

Without a collimator, we don't know the direction of the incoming photon

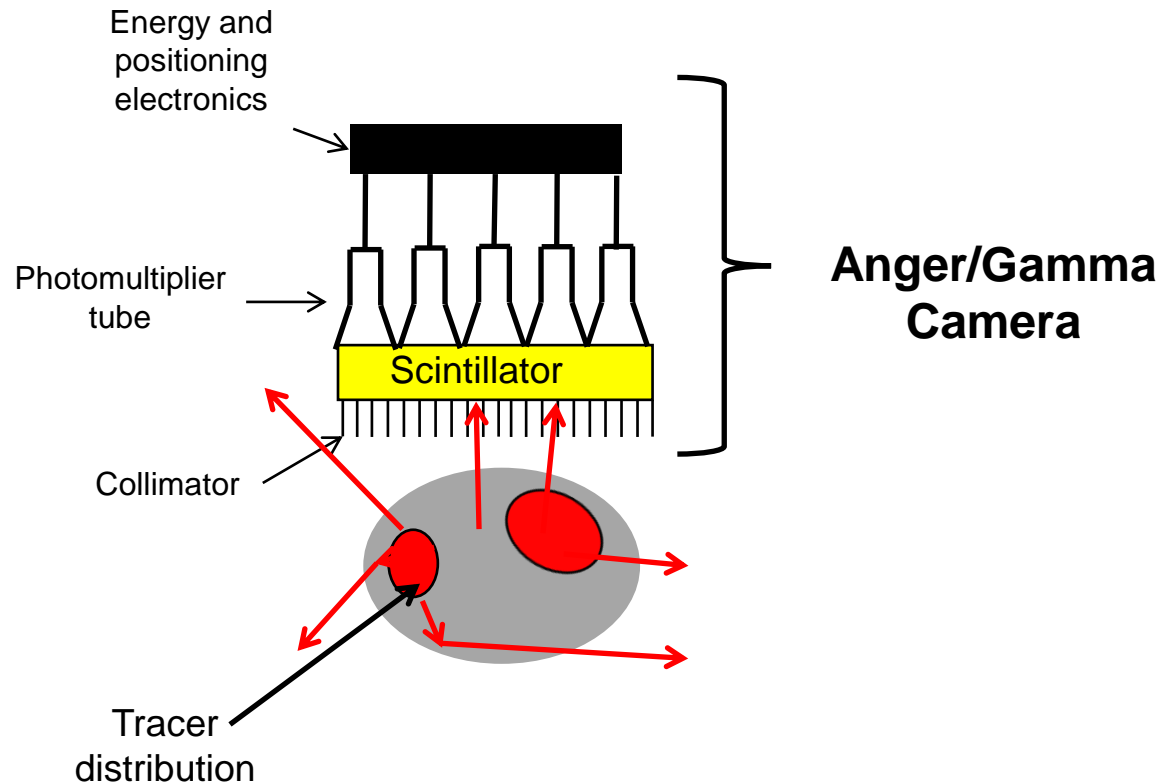
Absorptive Collimation



Requirements for collimator material:

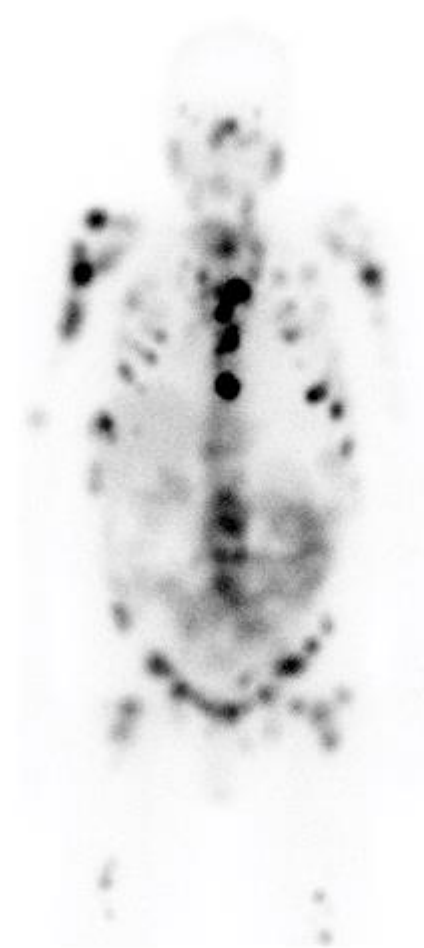
1. High attenuation coefficient μ at energy E of the emissions.
2. Should produce very few scatter photons.

Planar Imaging (2D)



- Gamma camera is used to detect gammas emitted from single-photon-emitting radionuclides

Planar Imaging (2D)

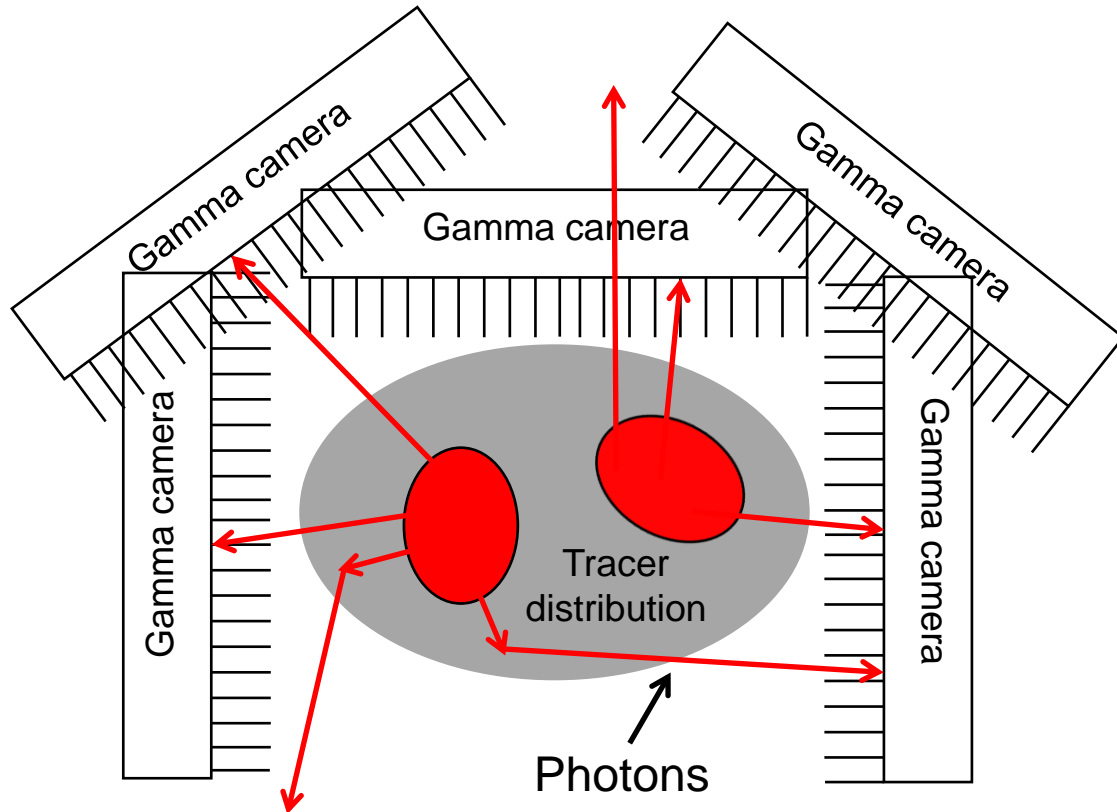


Anterior



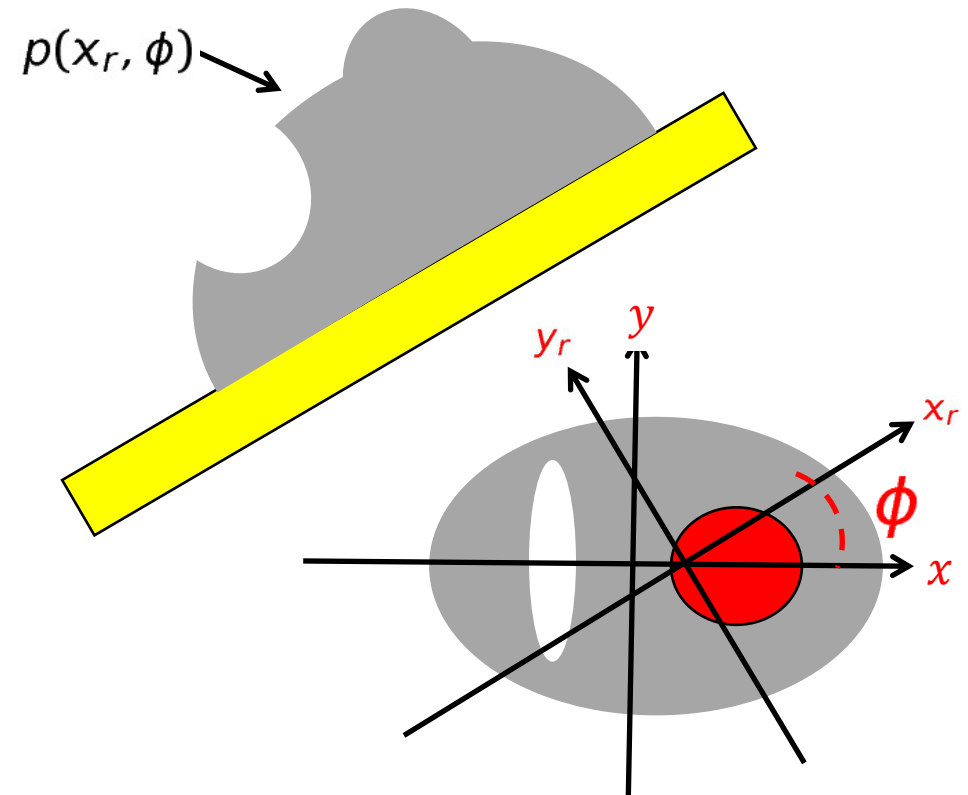
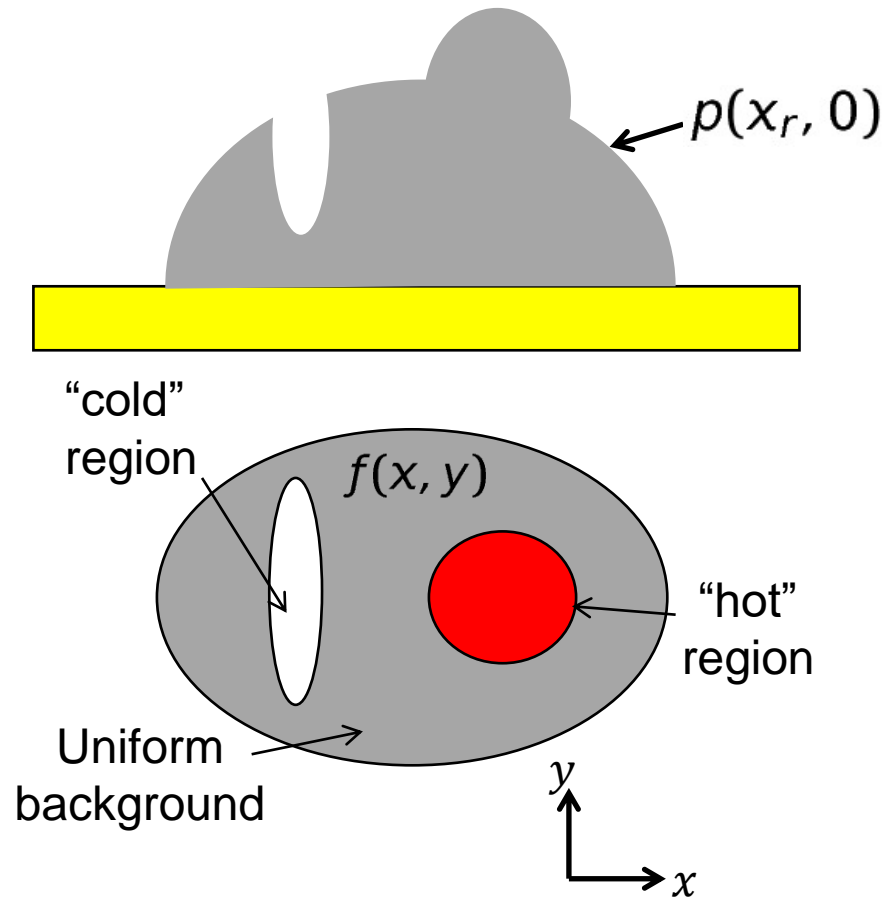
Posterior

Tomographic Imaging

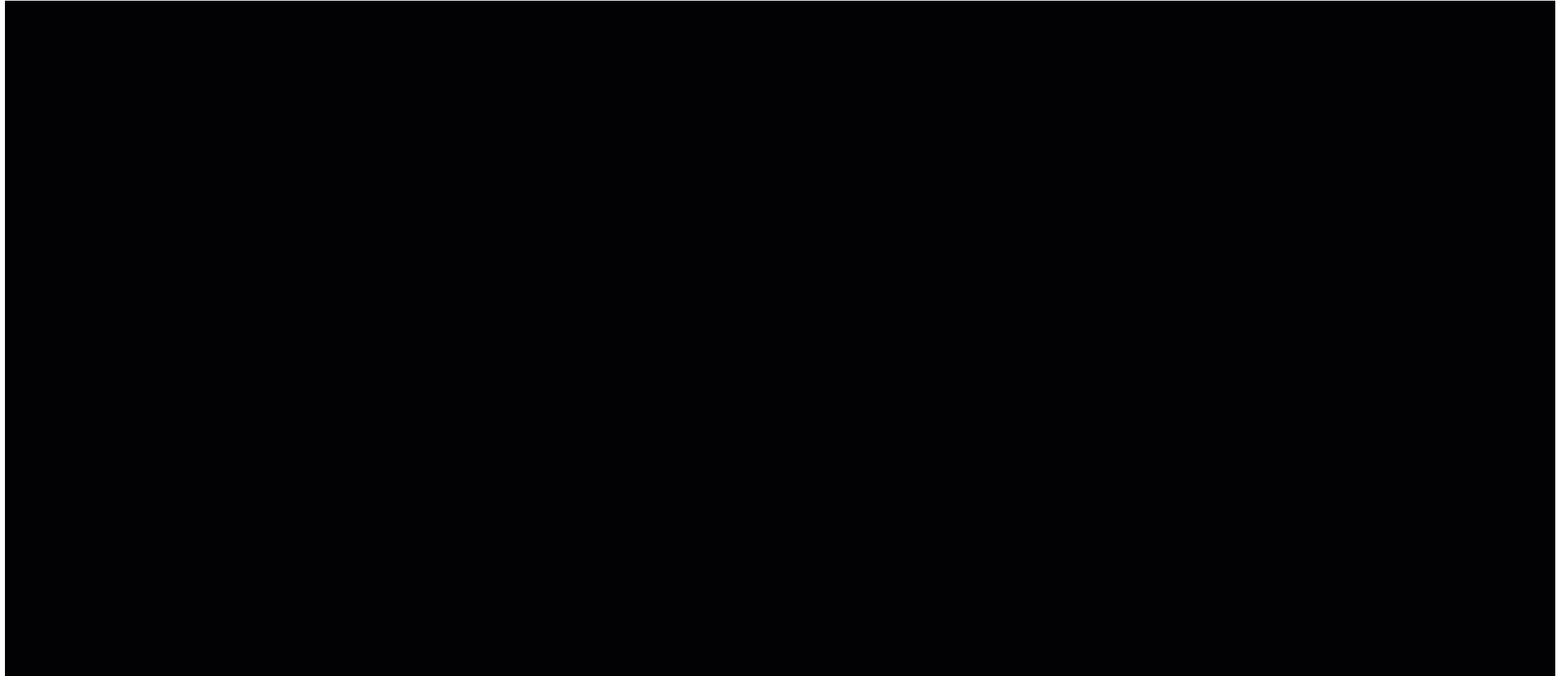


Camera is rotated to acquire *projections* at many angles enabling 3D imaging

Tomographic Imaging

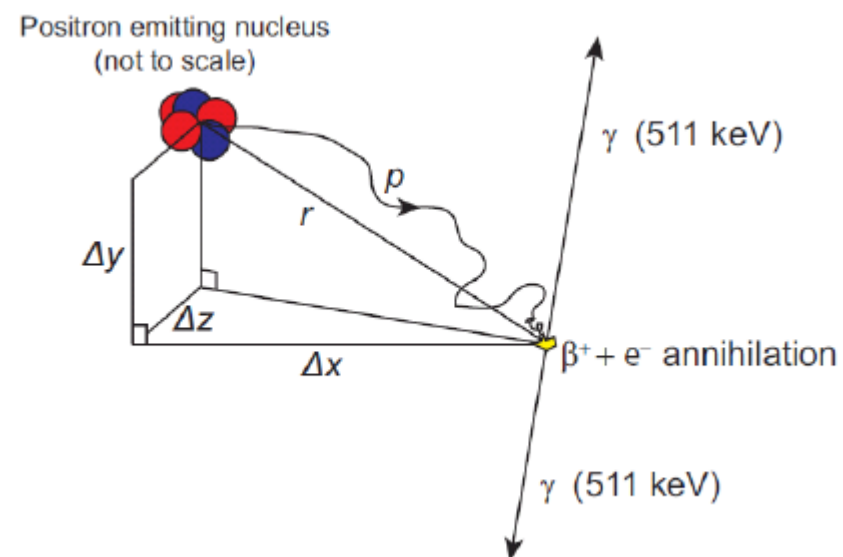


Tomographic Imaging SPECT

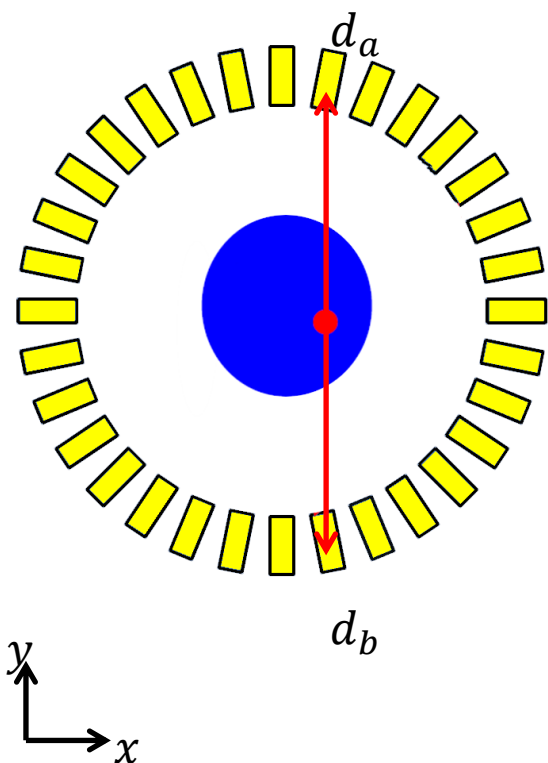


Positron Emission Tomography (PET)

- Nuclei undergoes β^+ decay
- The positron travels for a few mm and interacts with an electron in tissue
- An annihilation event occurs
- The annihilation produces two photons that travel in opposite directions

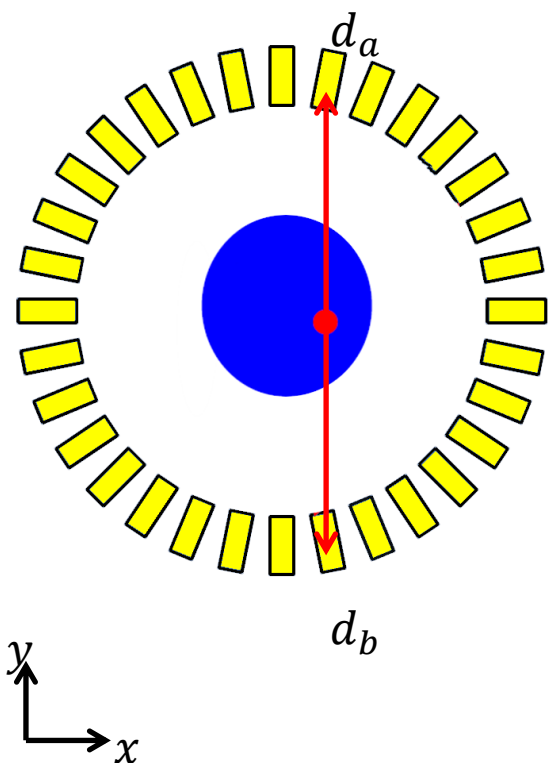


Positron Emission Tomography (PET)



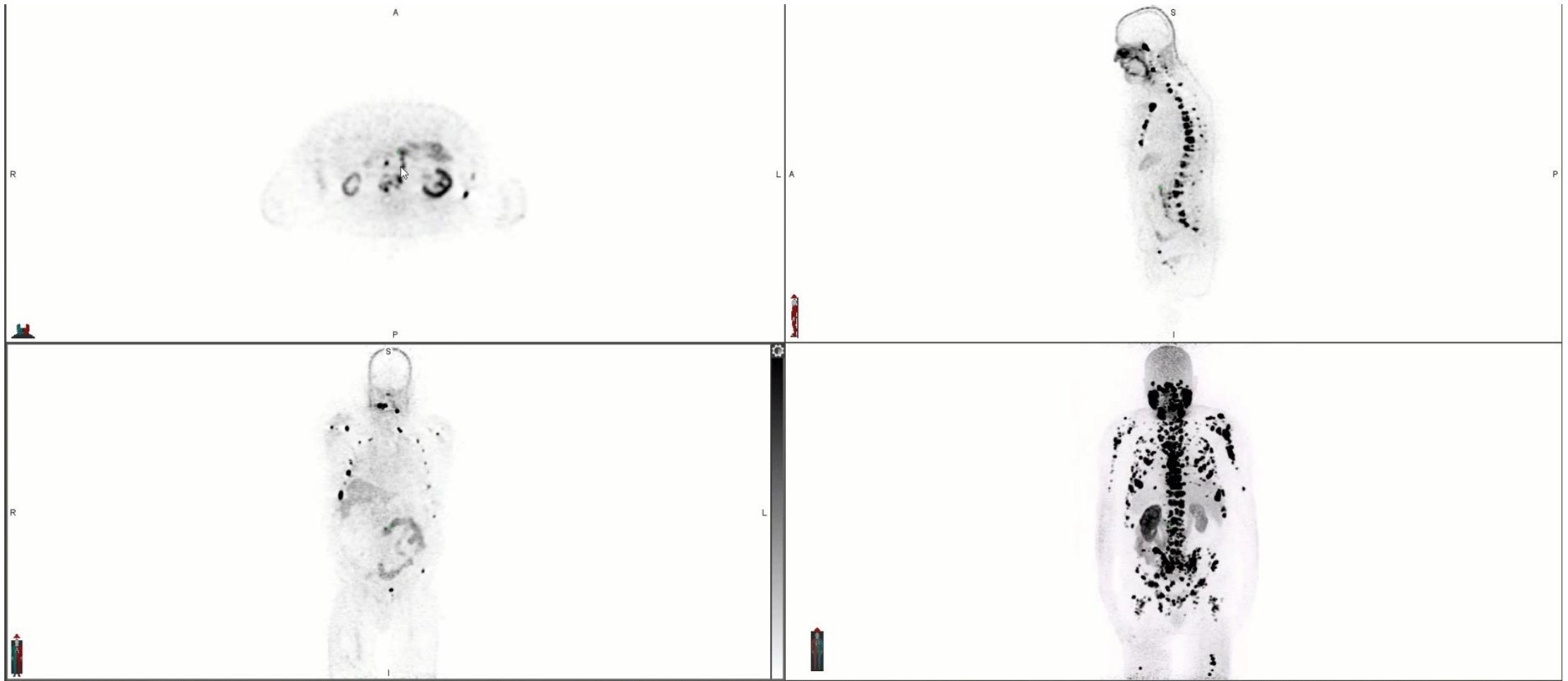
- Measure electrical pulses from pairs of detectors
- Timing pulses are triggered when the voltages exceed predefined thresholds

Positron Emission Tomography (PET)



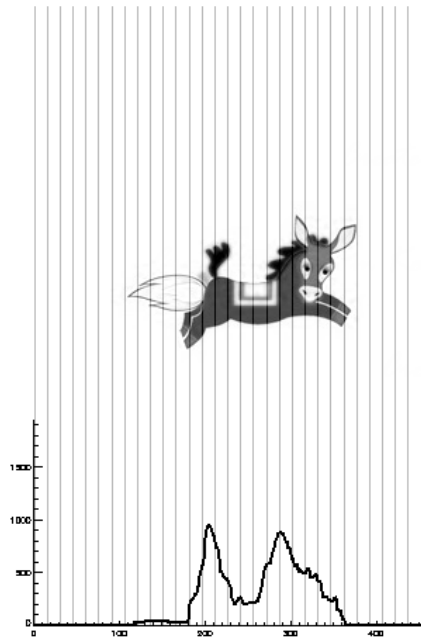
- The detector ring detects both of the 511 keV annihilation photons
- If the two events are detected within a small time difference (typically 6-12 ns) they are assumed to come from the same annihilation event
- The line of response passes through the point of annihilation
- No need for a collimator

Tomographic Imaging PET



Tomographic Imaging

(True) Emission Volume



intensity profile:



Sinogram (stored data)

Forward
Projection

angle
0

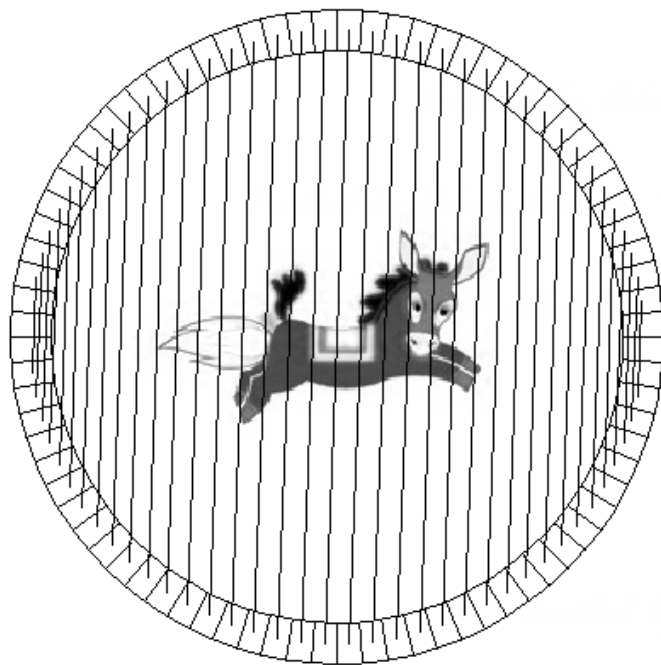
Theta (angle)

Rho (offset)

Created by Adam Leon Kesner, PhD, Medical Physicist University of Colorado, Anschutz Medical Campus, Aurora, Colorado, USA

Tomographic Imaging PET

Lines of response between PET detectors



Angle: 0°

Corresponding location in sinogram

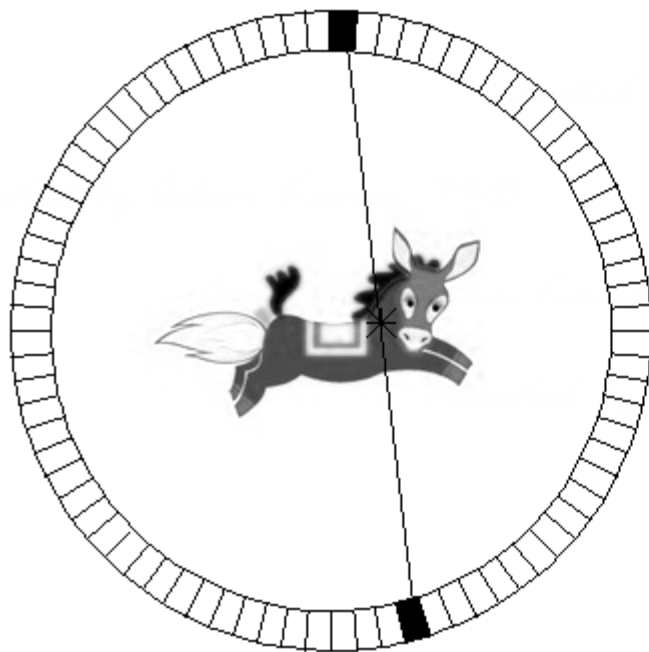


Rho (offset)

Created by Adam Leon Kesner, PhD, Medical Physicist University of Colorado, Anschutz Medical Campus, Aurora, Colorado, USA

Tomographic Imaging PET

Emission volume + PET detectors



Sinogram (histogram)



Theta (angle)

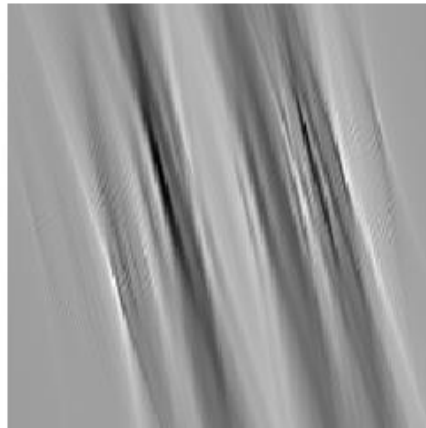
Rho (offset)

Events processed: 1

Created by Adam Leon Kesner, PhD, Medical Physicist University of Colorado, Anschutz Medical Campus, Aurora, Colorado

Image Reconstruction

Reconstructed image



Percent
backprojected
5 %
from selected
angles

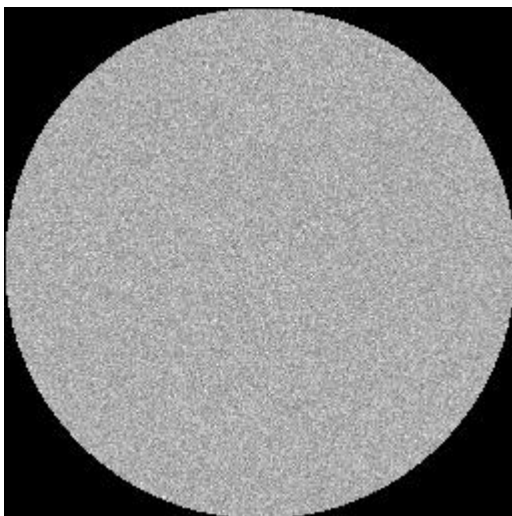
Sinogram



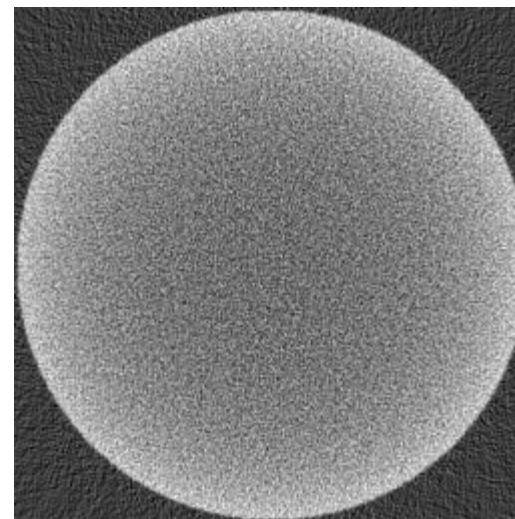
← Filtered Back Projection

Created by Adam Leon Kesner, PhD, Medical Physicist University of Colorado, Anschutz Medical Campus, Aurora, Colorado, USA

Anything wrong with this image?



$f(x, y)$

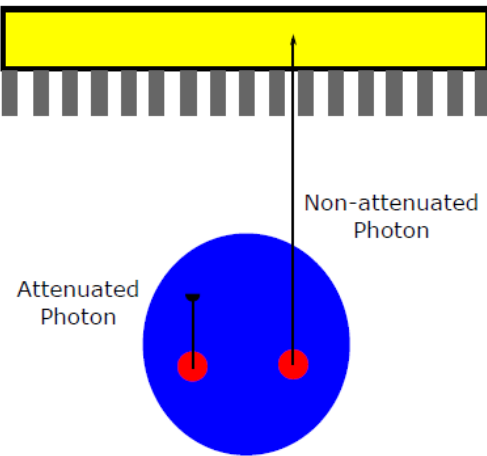


Reconstructed image

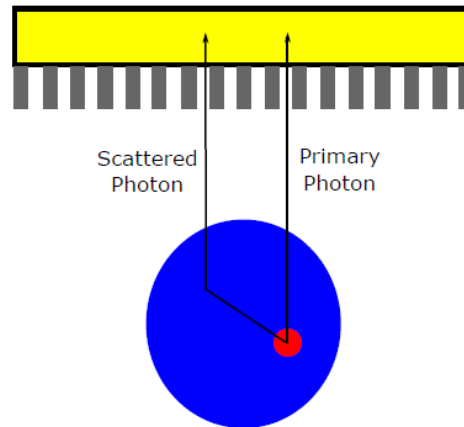
Disk with uniform activity concentration

Quantitative Imaging

Image Degrading Effects SPECT

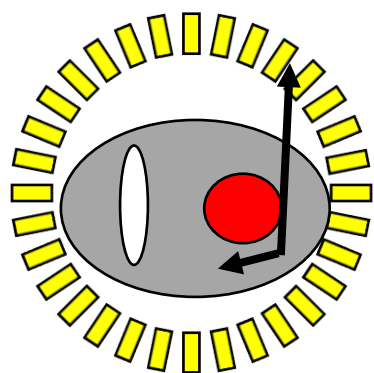


Attenuation

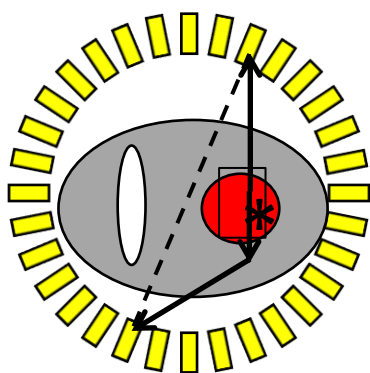


Scatter

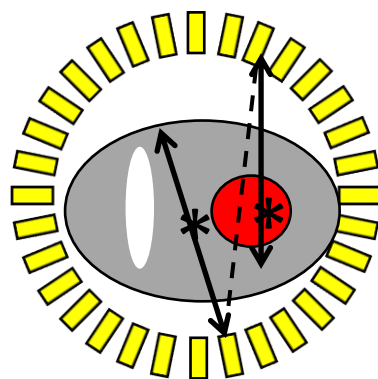
Image Degrading Effects PET



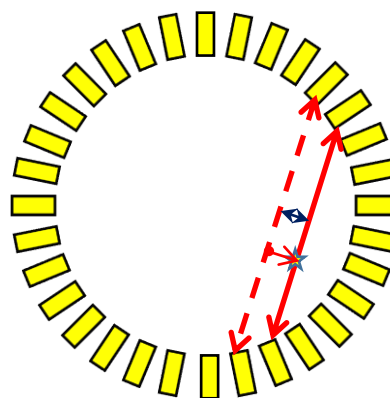
Attenuation



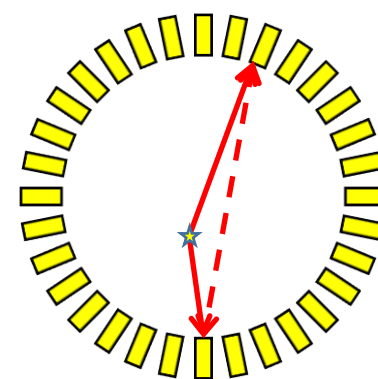
Scatter



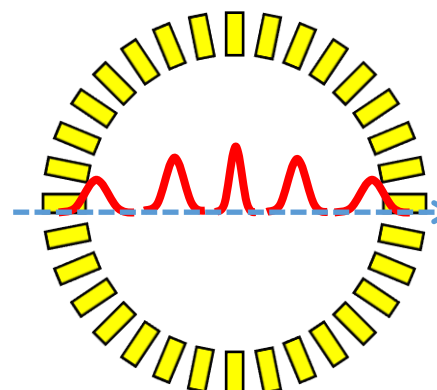
Randoms



Positron range

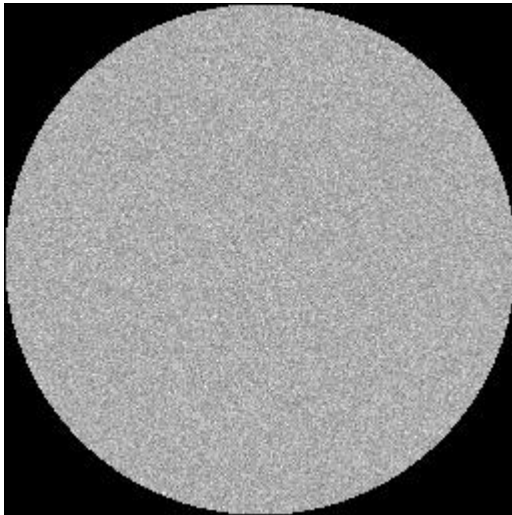


Positron
non-collinearity

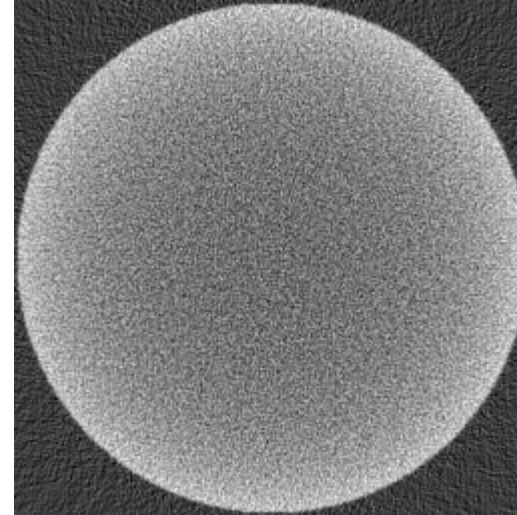


Depth of
interaction

But there's a problem...



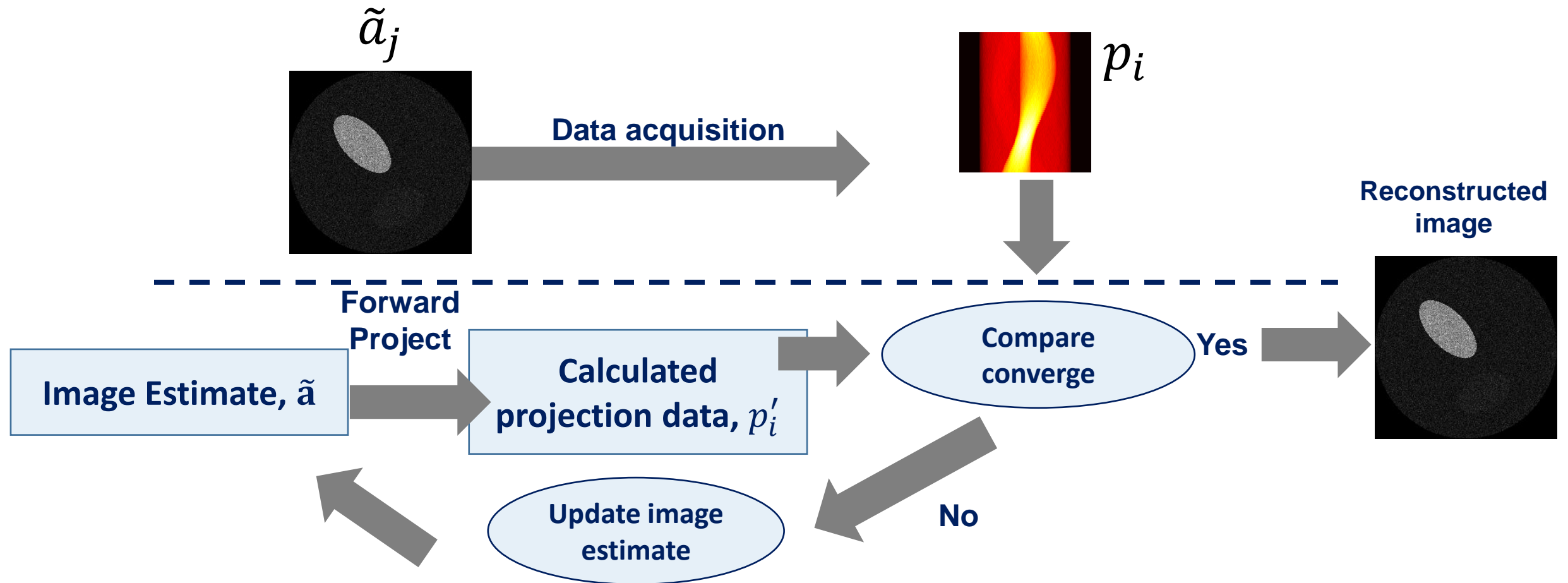
$f(x, y)$



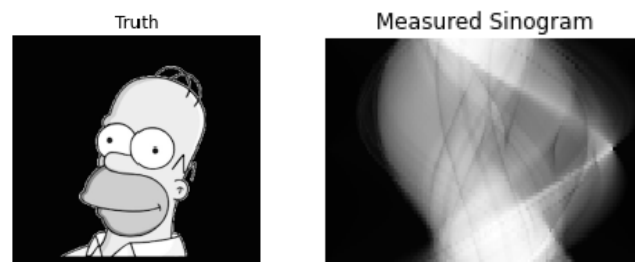
Reconstructed image

Disk with uniform activity concentration

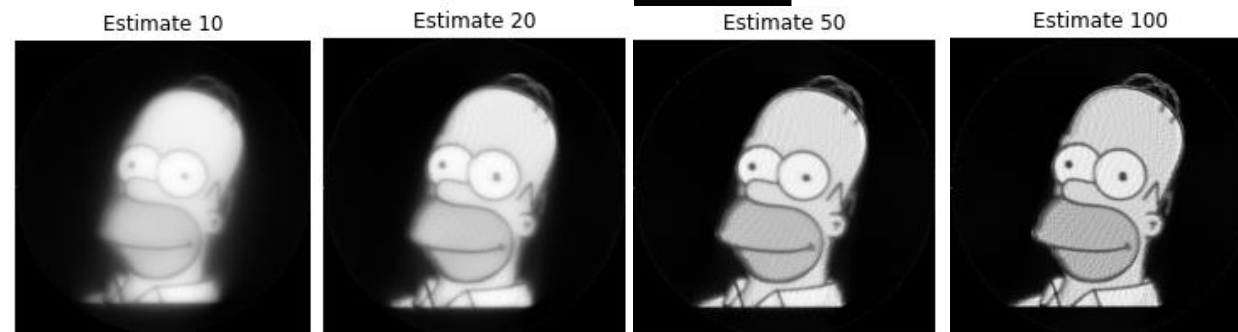
Iterative reconstruction algorithms



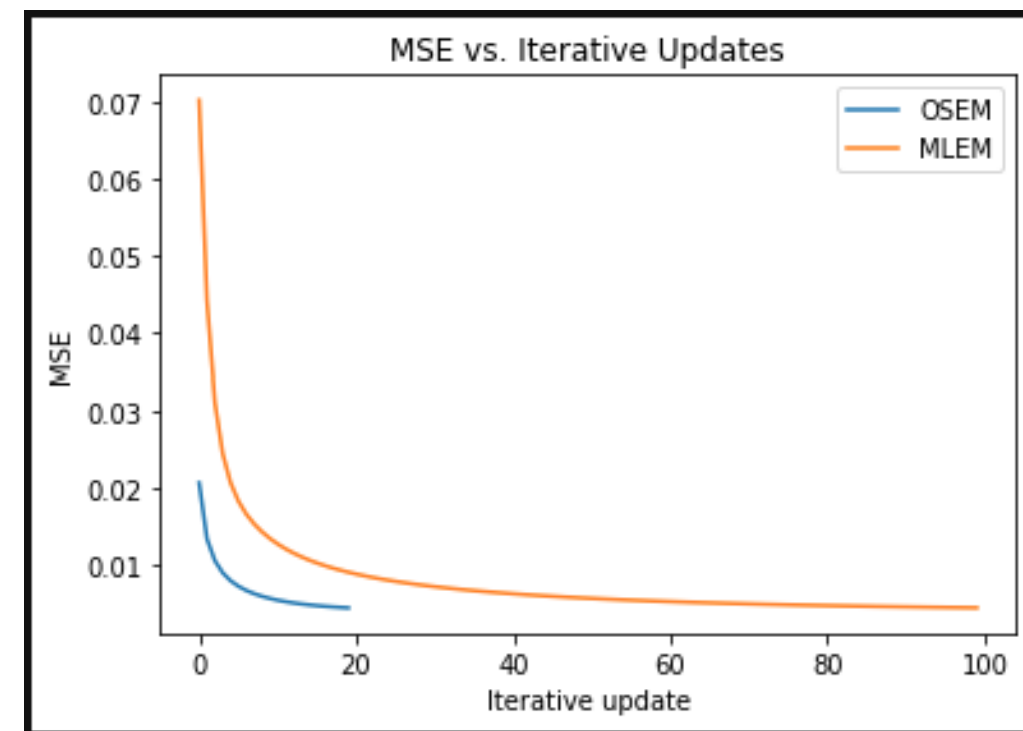
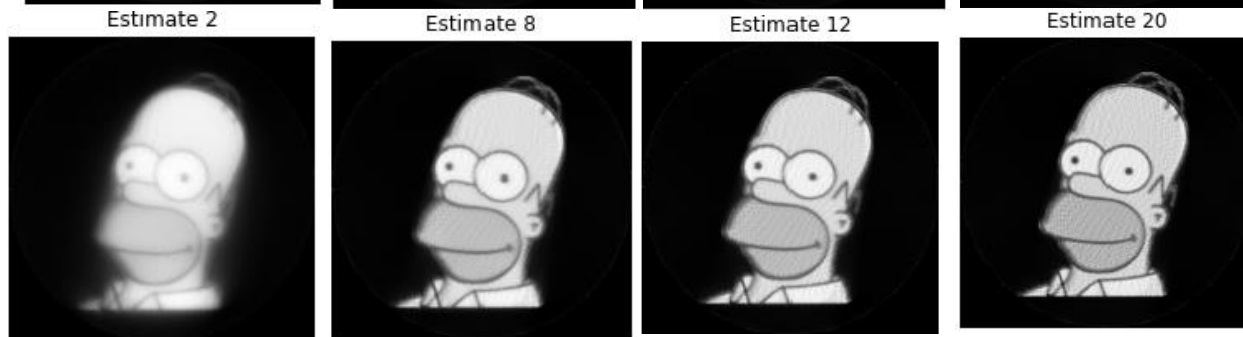
Iterative reconstruction algorithms



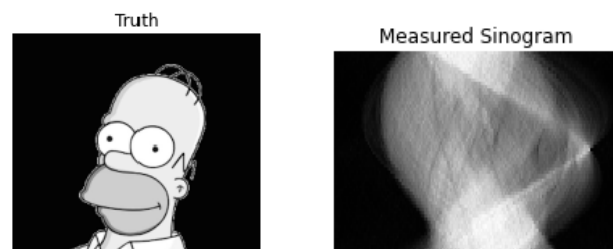
MLEM



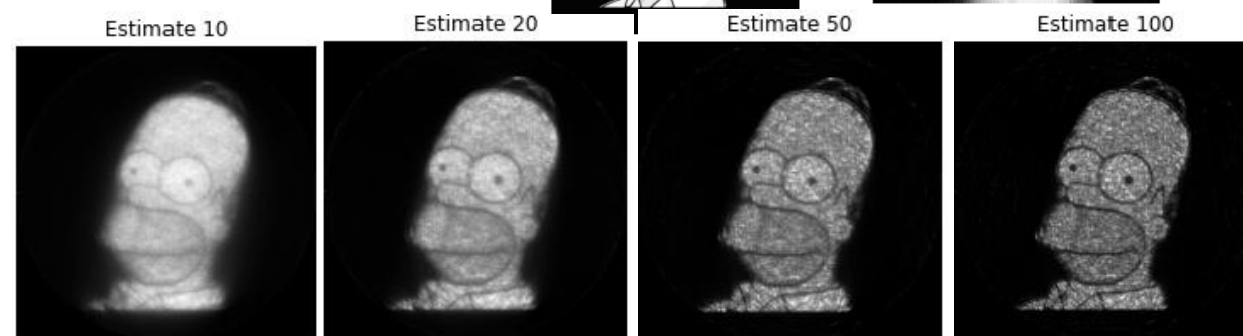
OSEM



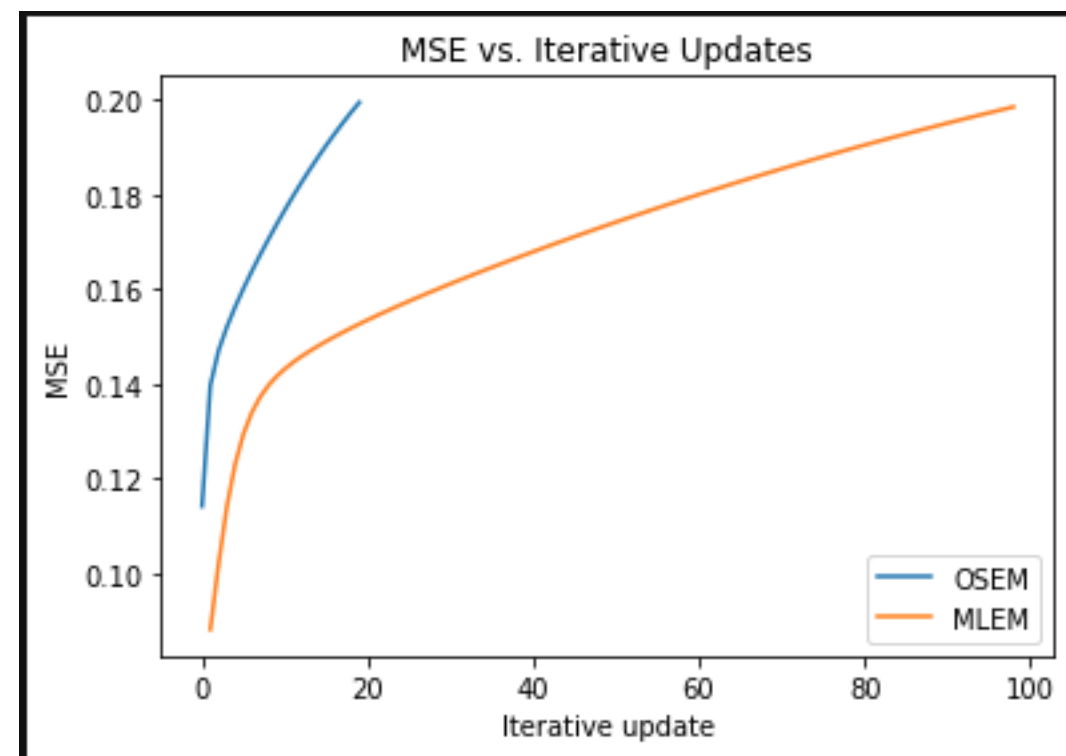
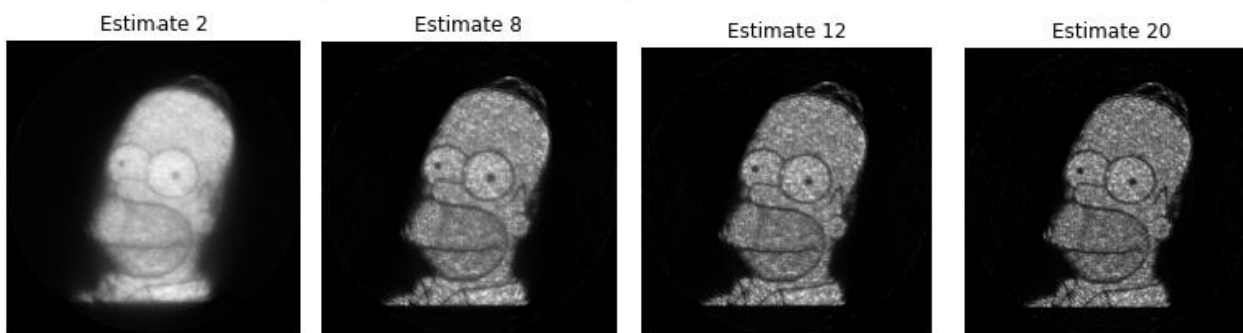
Iterative reconstruction algorithms



MLEM



OSEM



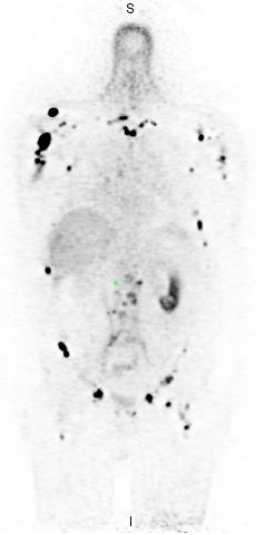
https://github.com/carluri/nucmed_physics_examples/tree/master/Lab2

Quantitative Imaging

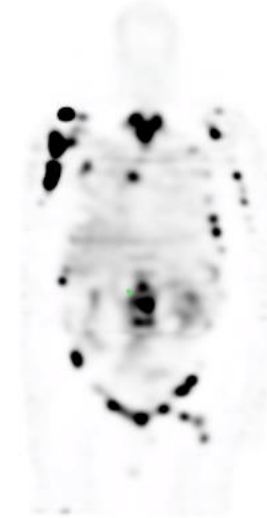
- If SPECT and PET images are **corrected for degrading effects**, then the image should contain information about the primary photons originating inside the patient
- **The images now have the correct number of photons in each pixel**
- However, a calibration factor is still required in order to convert the counts into activity or activity concentration values
- In order to obtain it, a scan of a perfectly known activity, with all the corrections is performed.

Quantitative Imaging

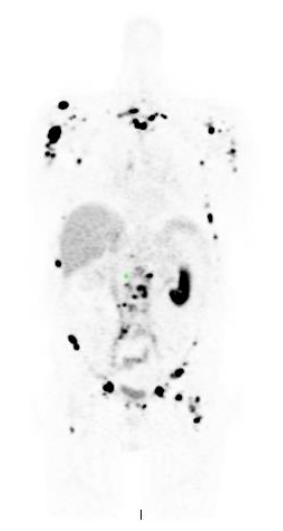
PET
No
Corrections



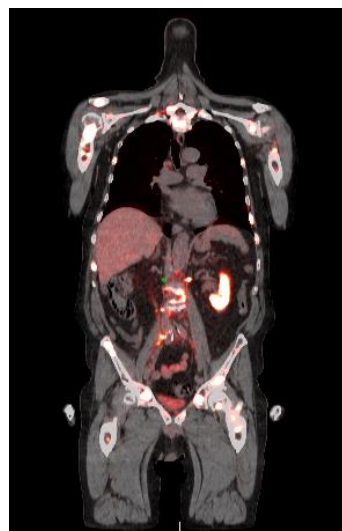
SPECT
No
Corrections



CT



PET Corrected



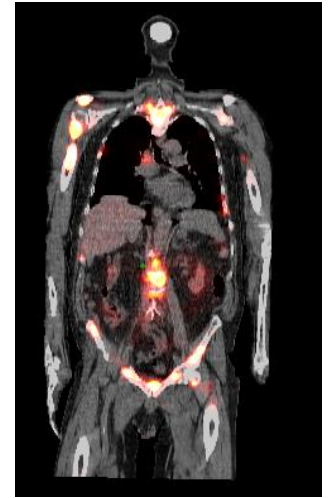
Fused Image



CT



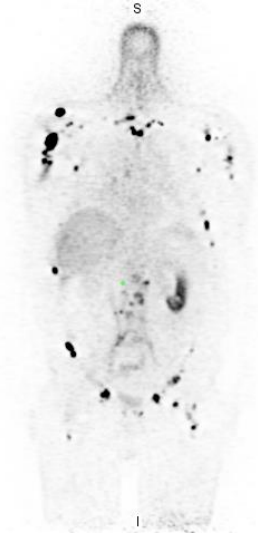
SPECT Corrected



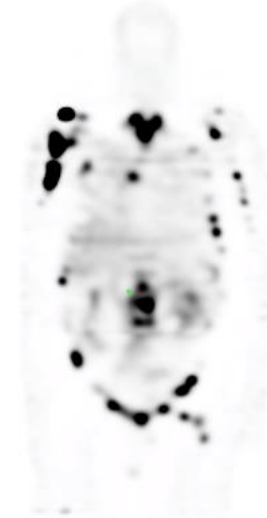
Fused Image

Quantitative Imaging

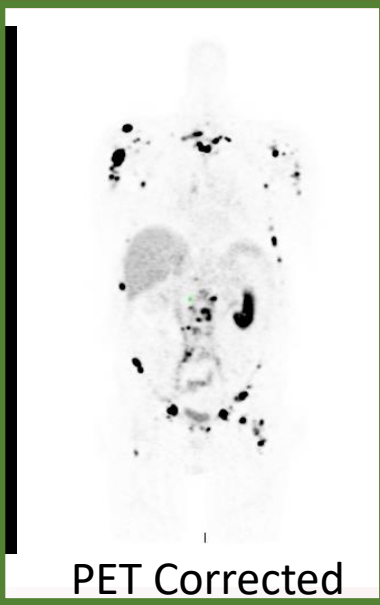
PET
No
Corrections



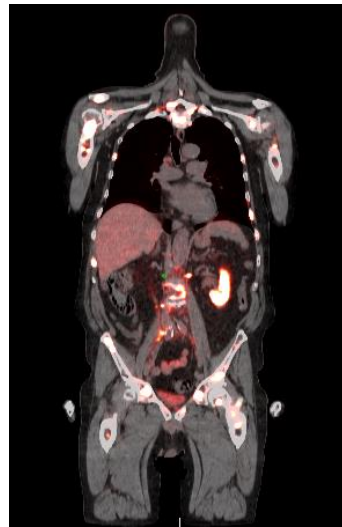
SPECT
No
Corrections



CT



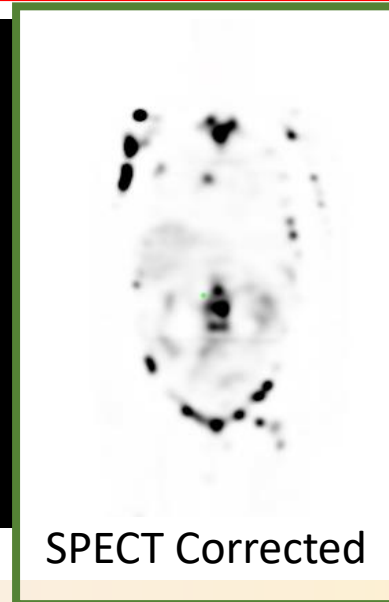
PET Corrected



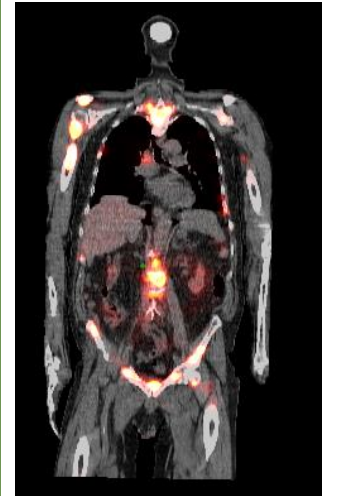
Fused Image



CT



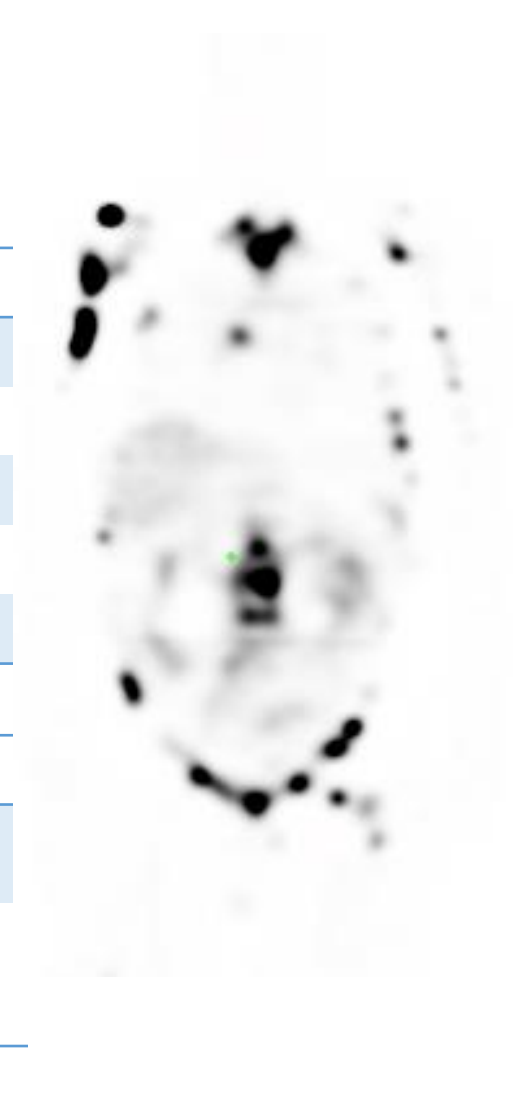
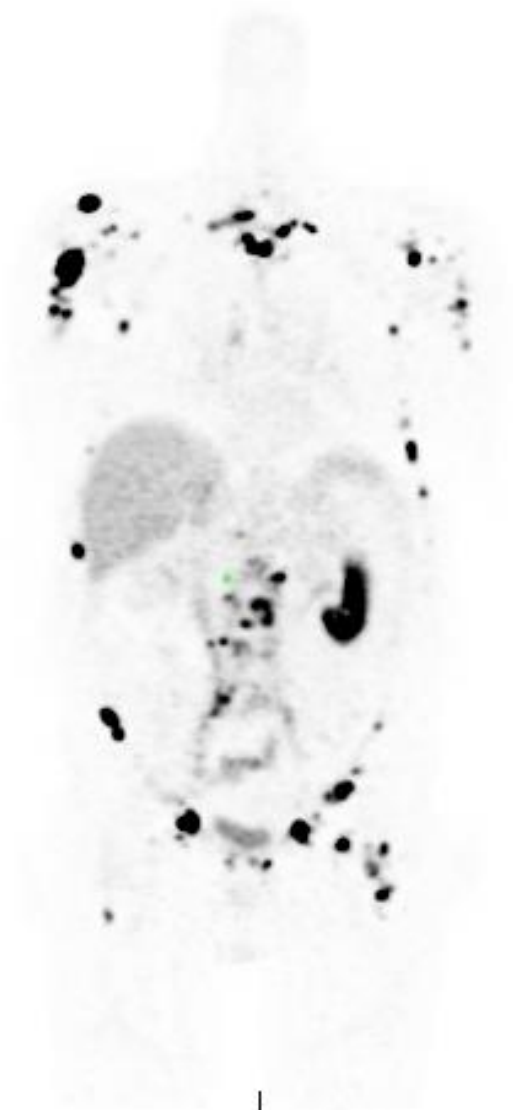
SPECT Corrected



Fused Image

Theranostics

PET and SPECT scans



PET	SPECT
[⁶⁸ Ga]Ga-PSMA-11	[¹⁷⁷ Lu]Lu-PSMA617

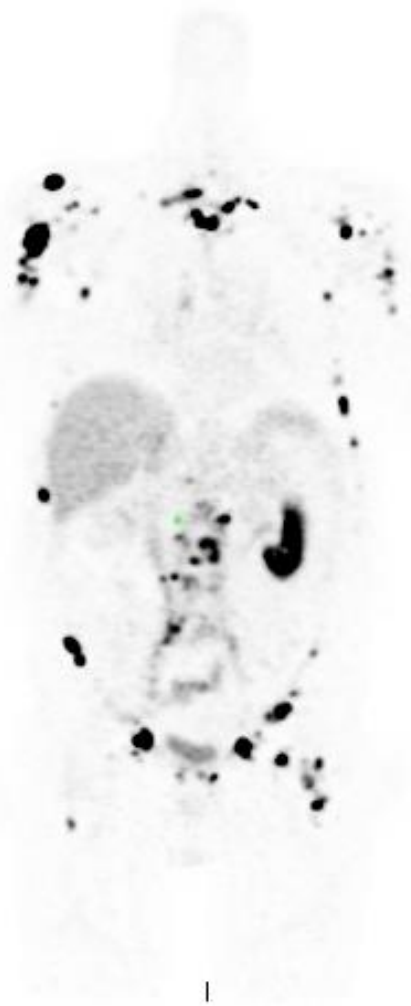
PET Isotopes (Positron Emitters)

Half-life	Rmean (mm)	Isotope
110 min	0.6	¹⁸ F
68 min	3.5	⁶⁸ Ga
20.4 min	1.2	¹¹ C
78.4 h	1.3	⁸⁹ Zr

Therapy Isotopes (Beta emitters)

Half-life	Rmean (mm)	Gamma emissions [keV]	Isotope
6.7 days	0.5	113 (6.2) 208 (10.3)	¹⁷⁷ Lu

PET and SPECT Scans



Diagnostic Scan

PET	SPECT
[⁶⁸ Ga]Ga-PSMA-11	[¹⁷⁷ Lu]Lu-PSMA617

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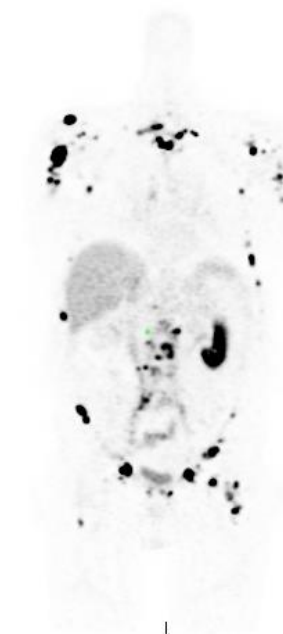
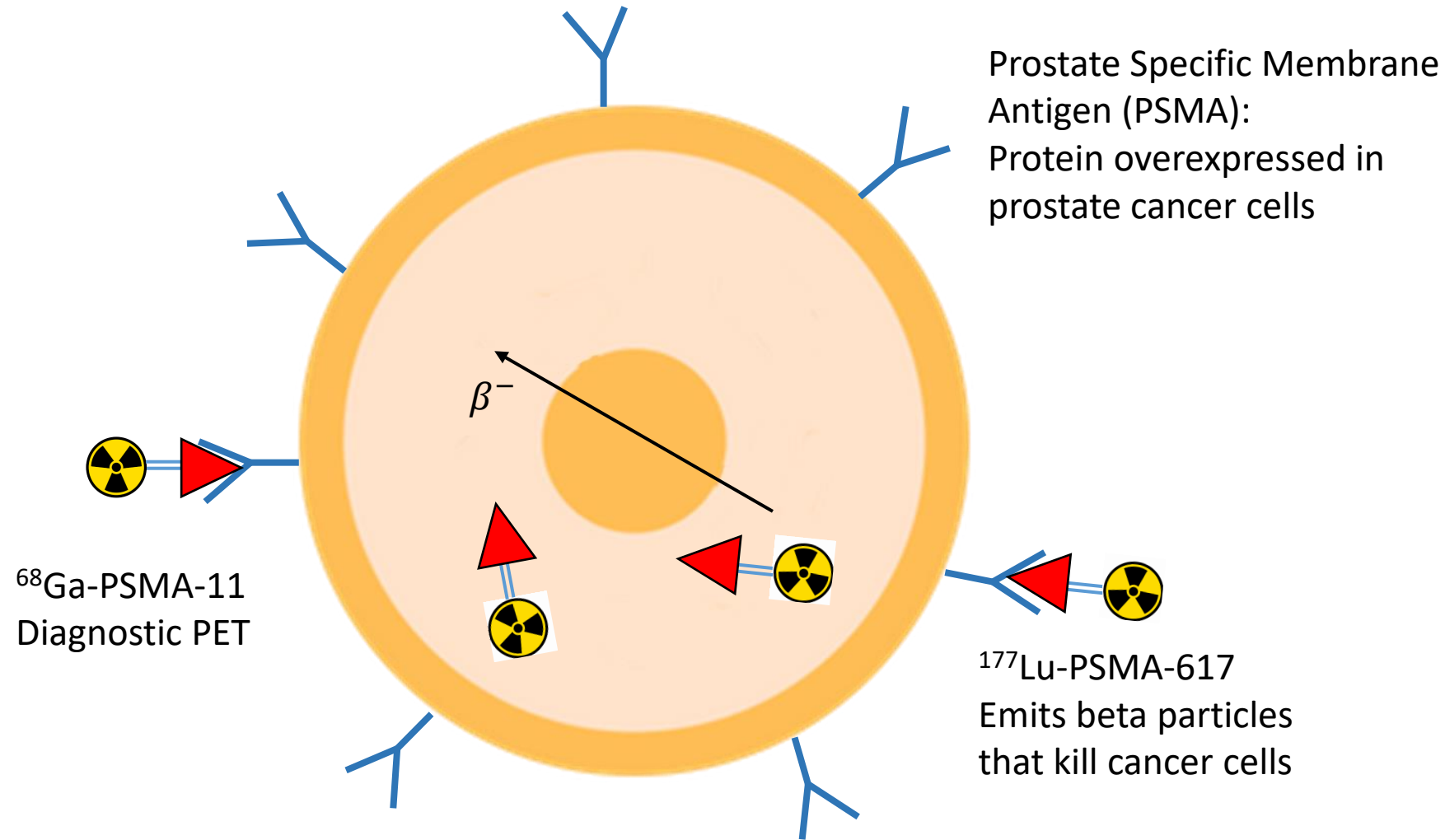
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Half-life	Rmean (mm)	Gamma emissions [keV]	Isotope
6.7 days	0.5	113 (6.2) 208 (10.3)	¹⁷⁷ Lu



Therapy Scan

Theranostics

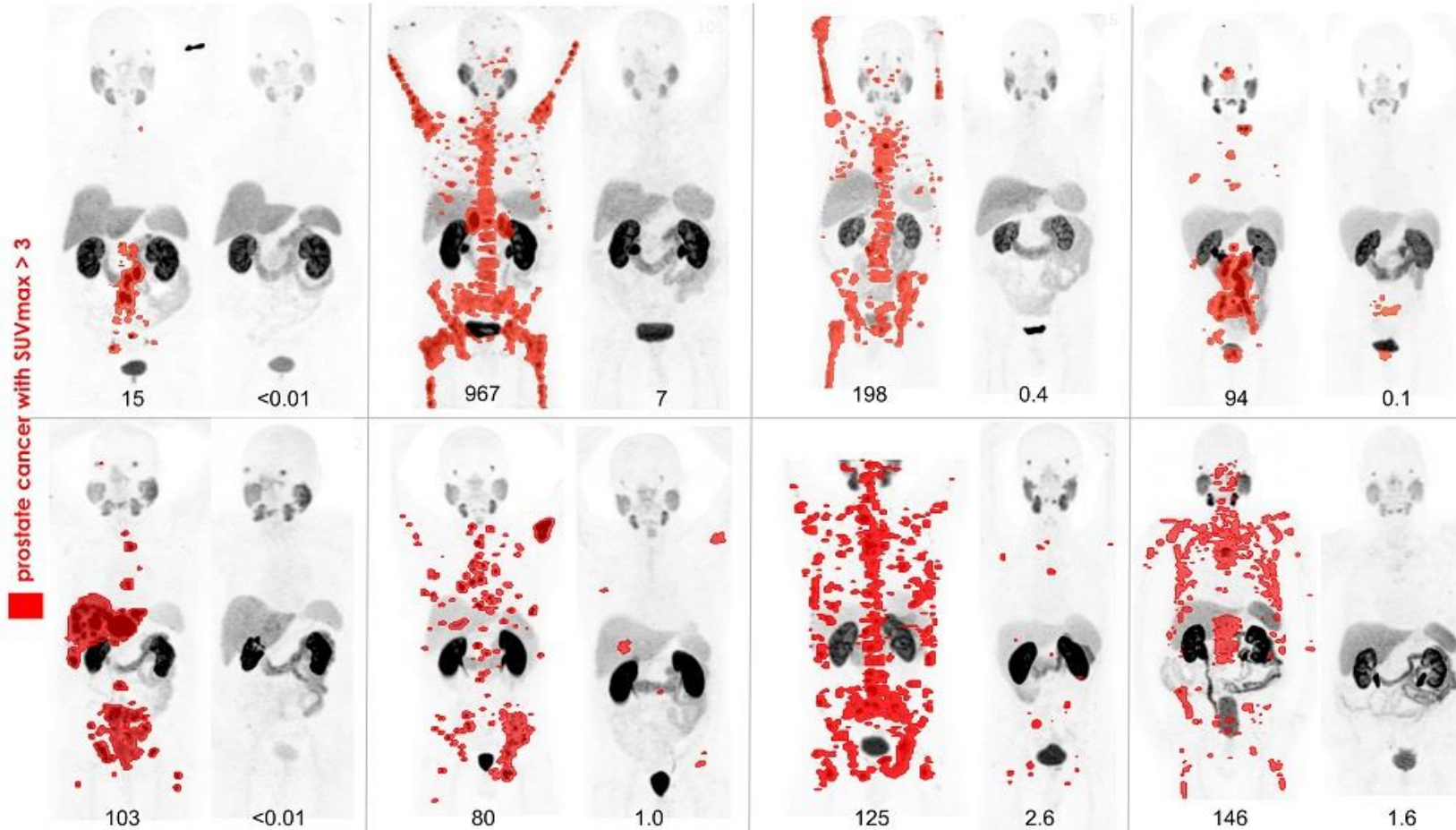


Diagnostic Scan



Therapy Scan

Theranostics - PSMA

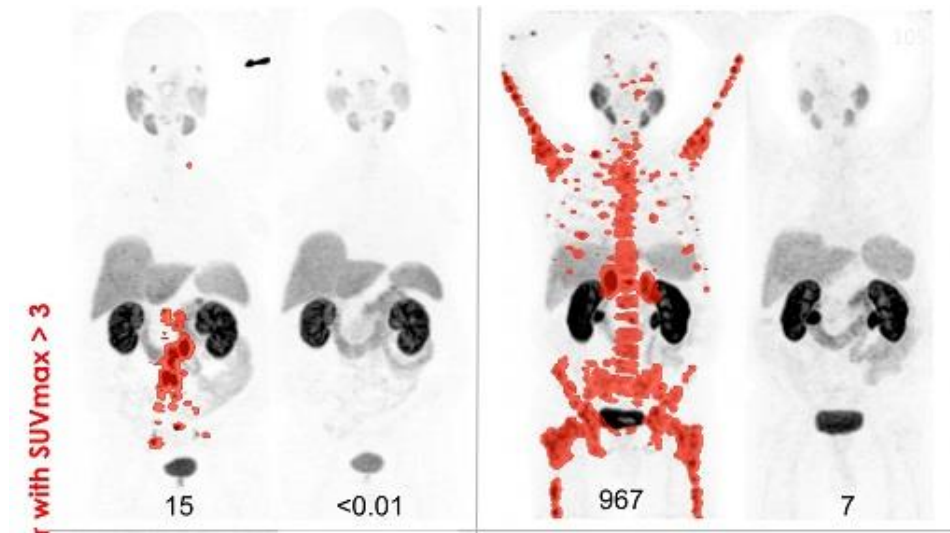


Hofman, et al. “[177Lu]-PSMA-617 Radionuclide Treatment in Patients with Metastatic Castration-Resistant Prostate Cancer (LuPSMA Trial): A Single-Centre, Single-Arm, Phase 2 Study.” *The Lancet Oncology* 19 (6): 825–33. 2018

Radiopharmaceutical Therapies (RPT)

Currently RPTs follow an empiric approach in which

- All patients are injected the same amount of radioactivity
- This approach limits the radiation toxicity to the population. Is possible that many patients are “under-treated”.
- Many of them can probably tolerate a higher radiation dose.

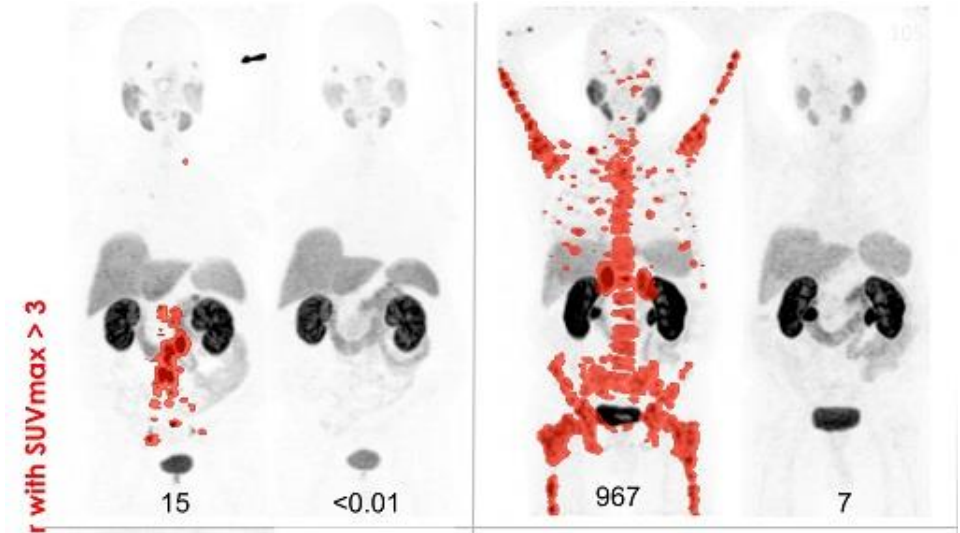


Personalized RPTs

If we perform personalized dose assessments (as is routine in external beam radiation therapy):

We could optimize treatment response by:

- Delivering maximum possible dose to tumors
- While sparing healthy organs by keeping dose levels below toxic.



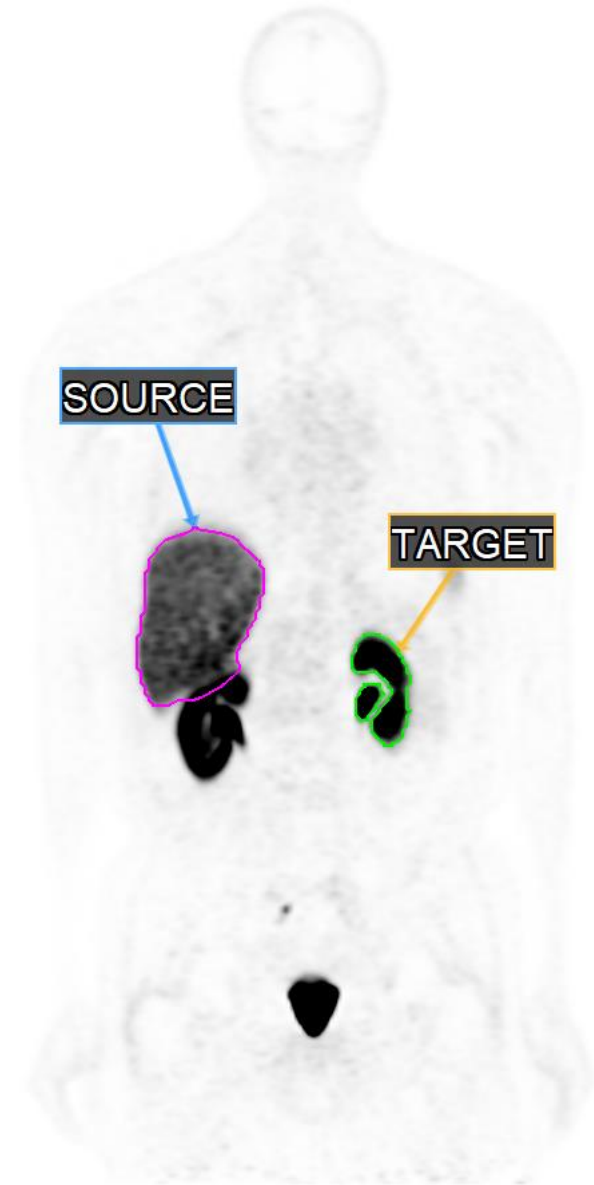
Internal Dosimetry

Dose

- Dose is defined as the energy deposited in a target region (e.g. organ or tumor) per unit mass

$$D = \frac{E}{M}$$

- SI units of absorbed dose is the joule per kilogram which has been named gray (1J/kg = 1 Gy).

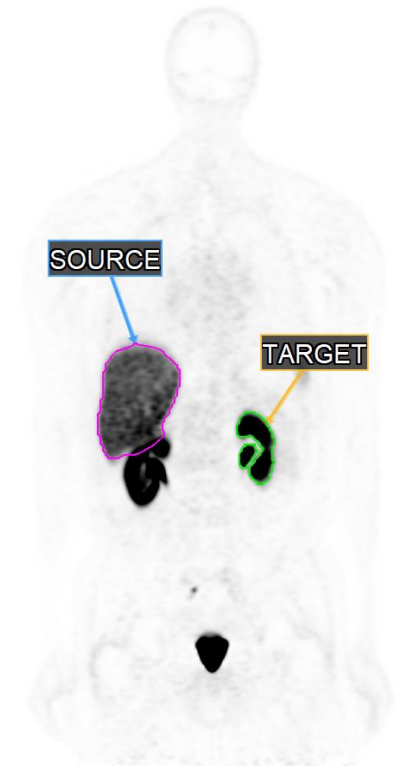


Internal Dosimetry

- To calculate dose in nuclear medicine procedures we need to have **information from the radioactive emissions of the radiopharmaceutical**

$$S(r_t \leftarrow r_s) = \sum_i \frac{n_i E_i \phi_i(r_t \leftarrow r_s)}{M_t}$$

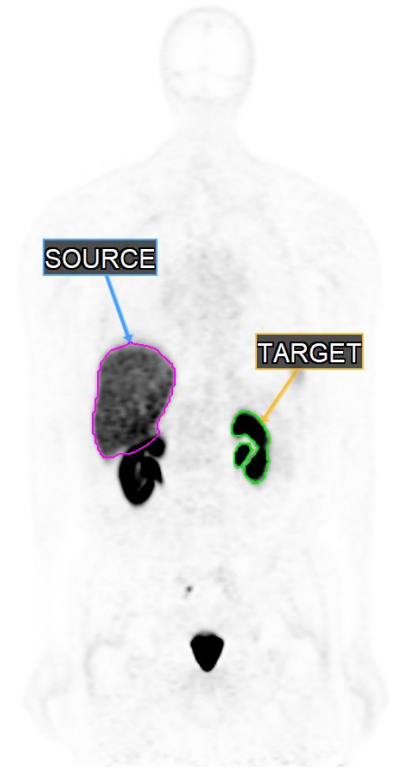
- Different decay modes of radionuclide of interest
 - Abundance (n_i) and energy (E_i) of each emission
- Absorbed fraction, $\phi_i(r_T \leftarrow r_S)$
- M_t mass of the target



Internal Dosimetry

$$S(r_t \leftarrow r_s) = \sum_i \frac{n_i E_i \phi_i(r_t \leftarrow r_s)}{M_t}$$

- The S-factor provides the dose absorbed by the target per decay in the source
- How many decays in total within the source?
- $A = \frac{dN}{dt} = A_0 e^{-\lambda t}$
- $N = \int A_0 e^{-\lambda t} dt$ } ???
- Biological processes (e.g. sweat, urine, etc)
- We need to incorporate a “biological half-life” that accounts for the rate at which the isotope is removed from the body

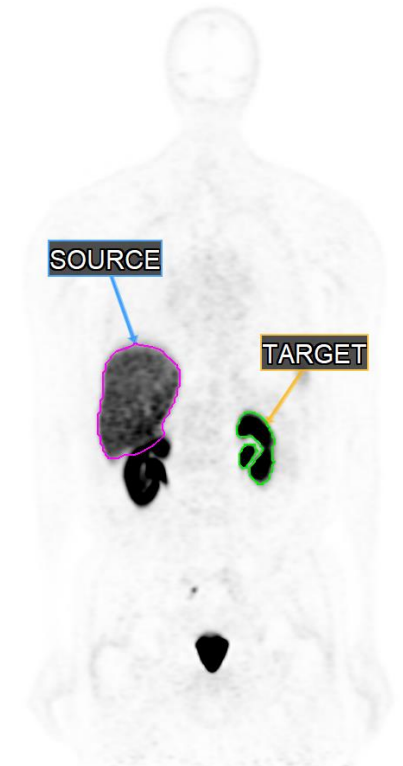


Internal Dosimetry

$$N = \int A_0 e^{-\lambda_{eff} t} dt = \tilde{A} \text{ Time Integrated Activity (TIA)}$$

$$S(r_t \leftarrow r_s) = \sum_i \frac{n_i E_i \phi_i(r_t \leftarrow r_s)}{M_t}$$

$$D(r_t \leftarrow r_s) = \tilde{A}(r_s) S(r_t \leftarrow r_s)$$

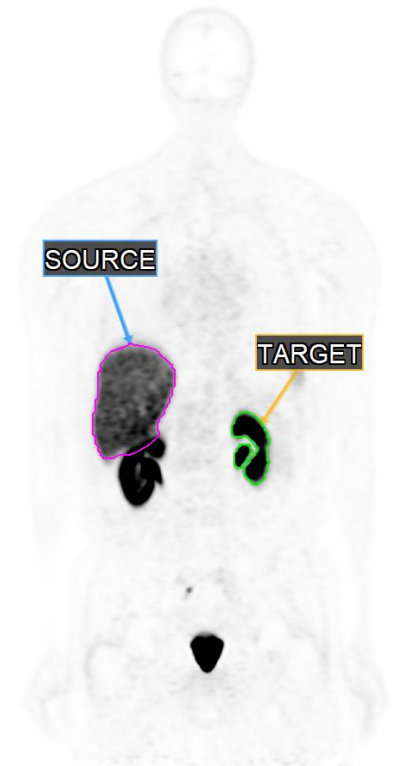


Internal Dosimetry

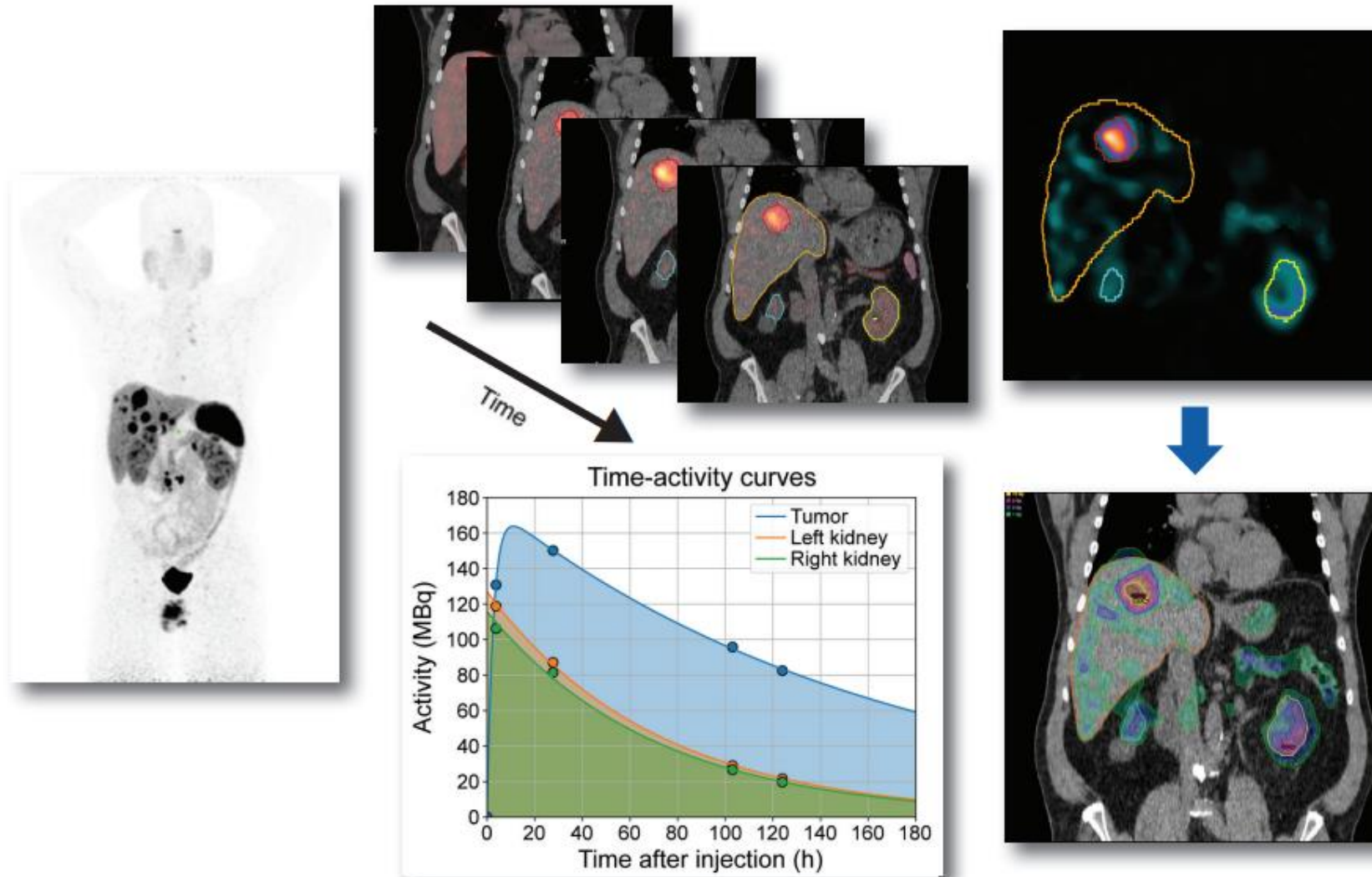
Biologic distribution data (we don't know it; we must measure it)

$$\bullet D(r_t \leftarrow r_s) = \tilde{A}(r_s) S(r_t \leftarrow r_s)$$

Details about physical properties of the radionuclide.
Method for combining biologic data with physical data to obtain Dose estimate



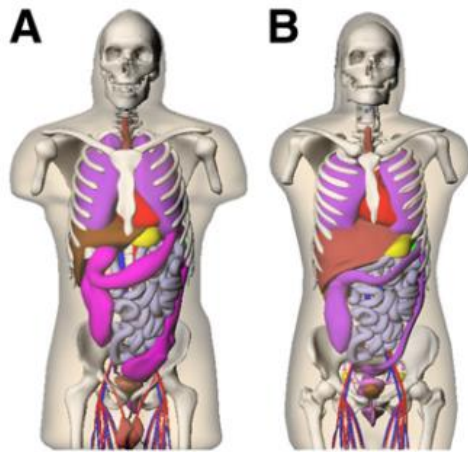
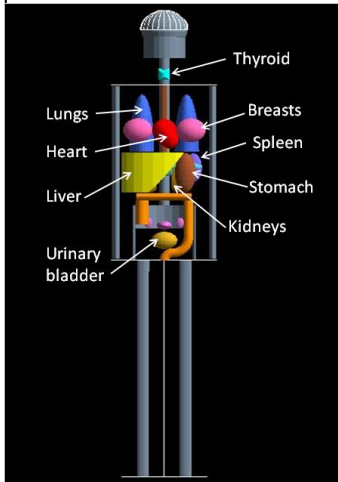
Dosimetry Workflow



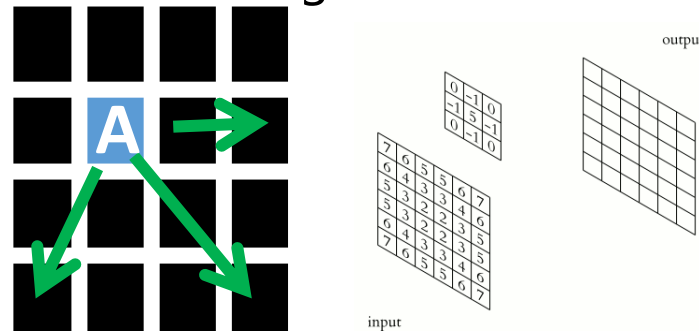
Dose Estimation Techniques

$$D(r_t \leftarrow r_s) = \tilde{A}(r_s) S(r_t \leftarrow r_s)$$

- Organ Level Approaches
 - Uses organ level S values
 - Dose to reference phantom



- Voxel S value approaches
 - Can deal with heterogeneous activity distributions
 - Precalculated dose distribution at voxel level
 - Homogeneous tissue



- Full Monte-Carlo
 - Uses patient's CT and 3D activity distribution
 - Simulates radiation transport of particles through media



Challenges

- Need quantitative image reconstruction
- Multiple measurements
 - When to measure?
 - How many times to measure?
 - Can the patient tolerate it?
 - Do we have enough resources?
- Image registration
 - We acquire images at different times. How do we know that one voxel in one image does correspond to another one in another image?

Challenges

- Image segmentation
 - How can we be sure that what we are segmenting is the truth?
 - What's the mass? What's the activity?
- How accurate is the used dosimetry model?
 - Can we report values with an uncertainty?
- Some of these procedures are very time consuming and need too many resources.
 - How can we simplify them? Make it easier to make the standard of care?

Challenges

- Despite a large number of clinical trials, considerable uncertainties still remain regarding the optimization of this therapeutic approach. The vital question still remains
- What is the dose-response relationship that could guide us in planning personalized treatments?

Our Vision and Efforts

Standardizing Dosimetry Protocols

Vol. 62 • Suppl. 3 • December 2021

IMPACT FACTOR
10.057

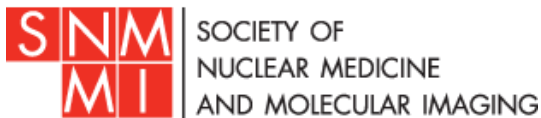
#1 NUCLEAR MEDICINE, MOLECULAR IMAGING AND MOLECULAR RADIOTHERAPY JOURNAL

JNM
The Journal of Nuclear Medicine

Radiopharmaceutical Dosimetry for Cancer Therapy: From Theory to Practice
Guest Editors: Richard L. Wahl, MD and John Sunderland, PhD

Illustration of steps in patient-specific dosimetry including pre-therapy PET quantitative SPECT at 4 timepoints with segmentation, integration of time-activity curves to form the time-integrated activity image, and the resulting dose map.

SNM Value Initiative
SOCIETY OF NUCLEAR MEDICINE & MOLECULAR IMAGING



JNM Supplement Addresses Radiopharmaceutical Dosimetry for Cancer Therapy - SNMMI

An International Study of Factors Affecting Variability of Dosimetry Calculations, Part 1: Design and Early Results of the SNMMI Dosimetry Challenge

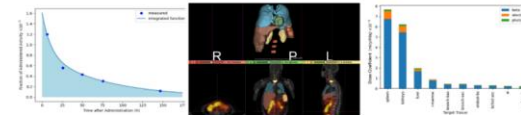
Carlos Uribe^{1,2}, Avery Peterson³, Benjamin Van³, Roberto Fedrigo⁴, Jake Carlson⁵, John Sunderland⁶, Eric Frey^{7,8}, and Yuni K. Dewaraja⁸

¹Functional Imaging, BC Cancer, Vancouver, British Columbia, Canada; ²Department of Radiology, University of British Columbia, Vancouver, British Columbia, Canada; ³Department of Radiology, University of Michigan Medical School, Ann Arbor, Michigan; ⁴Department of Integrative Oncology, BC Cancer Research Institute, Vancouver, British Columbia, Canada; ⁵U-M Library, University of Michigan, Ann Arbor, Michigan; ⁶Department of Radiology, University of Iowa, Iowa City, Iowa; ⁷Radiological Physics Division, Johns Hopkins University, Baltimore, Maryland; and ⁸Rapid, LLC, Baltimore, Maryland

Lu-177 Dosimetry Challenge 2021

Welcome to the SNMMI Lu-177 Dosimetry Challenge 2021

The SNMMI Dosimetry Task Force has the primary goal of advancing the use of dosimetry in radiopharmaceutical therapy (RPT). It has identified the need for harmonization of dosimetry methods as an area of focus. Although efforts to harmonize and standardize internal dosimetry calculations have been made, there has been a lack of large-scale studies to which to justify recommendations. The Task Force is thus soliciting members of the nuclear medicine community to contribute to the 177Lu Dosimetry Challenge.



What? Community science effort to provide data needed for harmonization and standardization of dosimetry methods for Radiopharmaceutical Therapy.

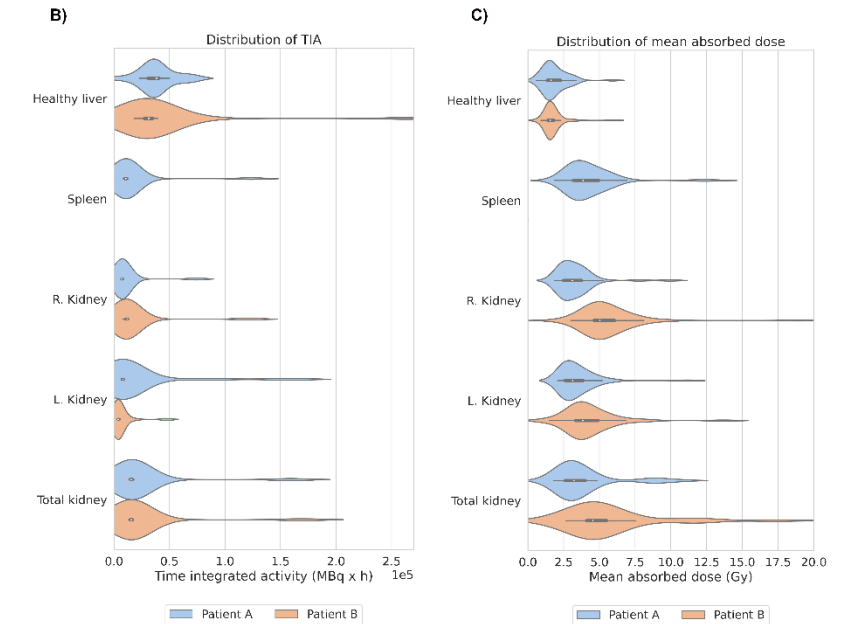
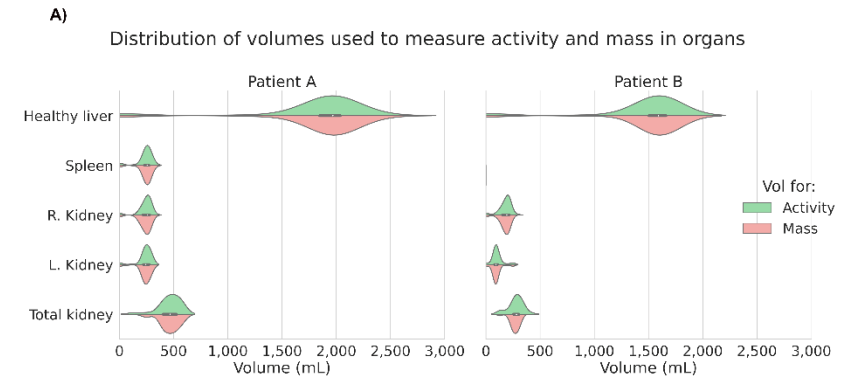
I WANT TO PARTICIPATE
REGISTER HERE

GOT QUESTIONS?
CONTACT US

Lu-177 DOSIMETRY CHALLENGE DETAILS AND INSTRUCTIONS

ACCESS

[Dosimetry Challenge \(snmmi.org\)](https://snmmi.org)

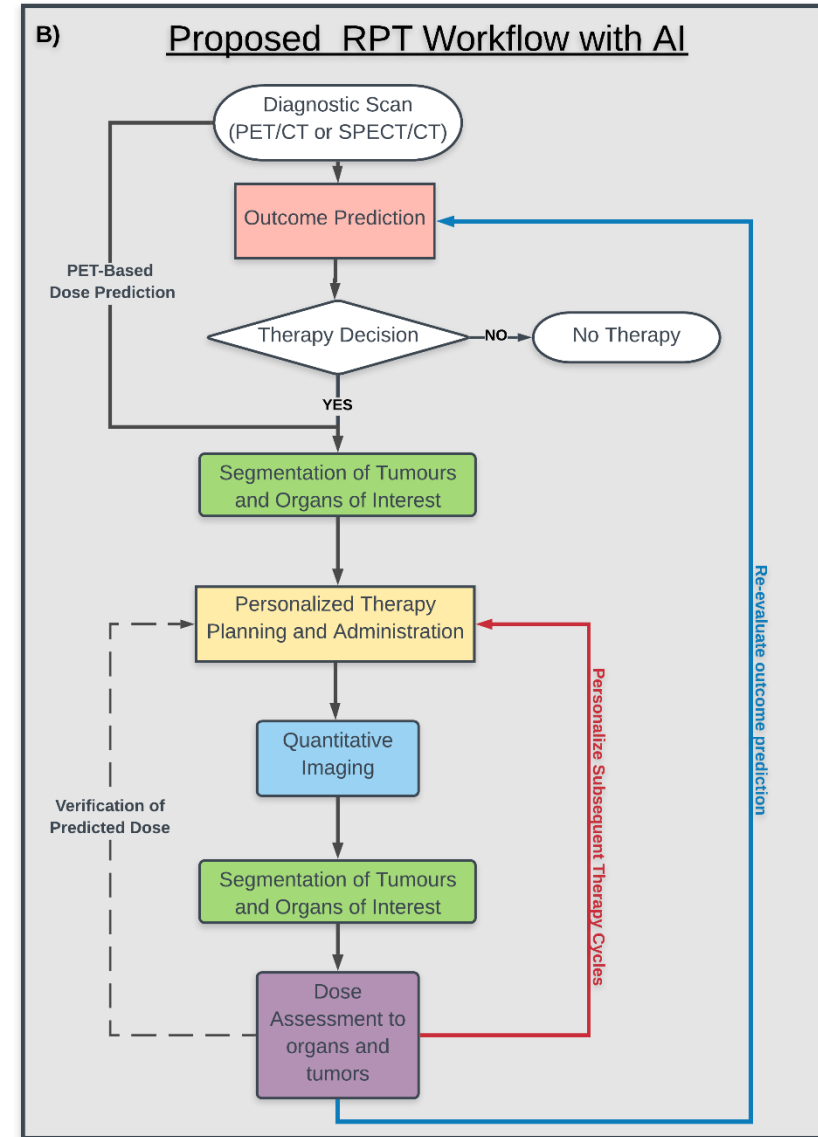


AI is a powerful tool

Role of Artificial Intelligence in Theranostics: Toward Routine Personalized Radiopharmaceutical Therapies

Julia Brosch-Lenz, PhD^a, Fereshteh Yousefirizi, PhD^a,
 Katherine Zukotynski, MD, PhD, FRCPC^b,
 Jean-Mathieu Beauregard, MD, MSc, FRCPC^{c,d}, Vincent Gaudet, PhD, PEng^e,
 Babak Saboury, MD, MPH, DABR, DABNM^{f,g,h},
 Arman Rahmim, PhD, DABSNM^{a,i,j}, Carlos Uribe, PhD, MCCPM^{i,k,*}

Brosch-Lenz, J. *et al.* *PET Clinics* **16**, 627–641 (2021).



Quantitative Imaging PSMA

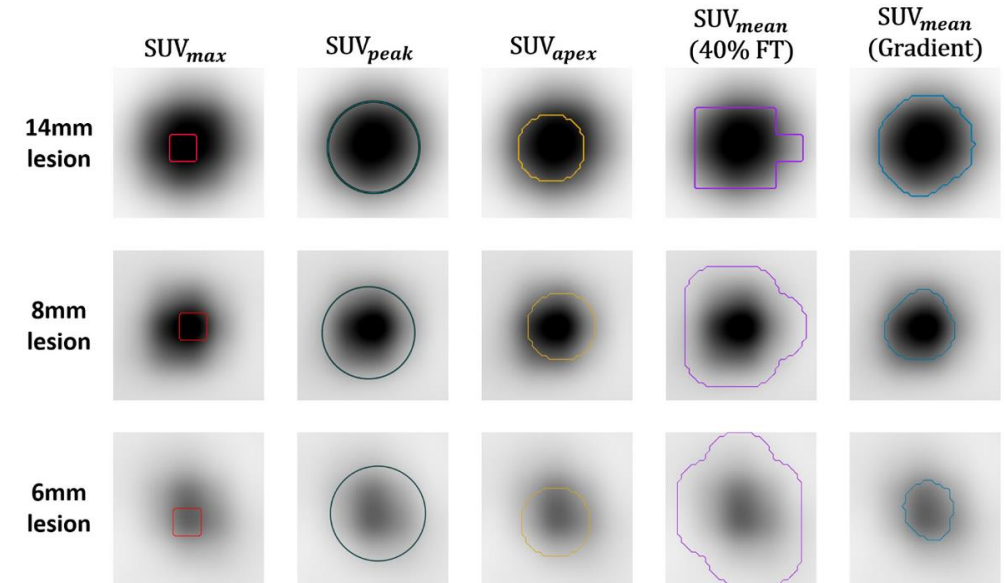
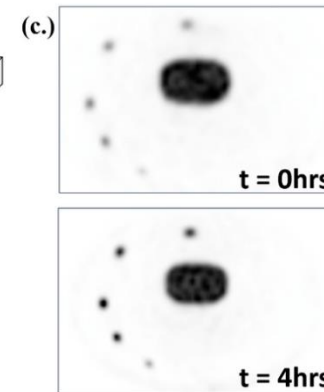
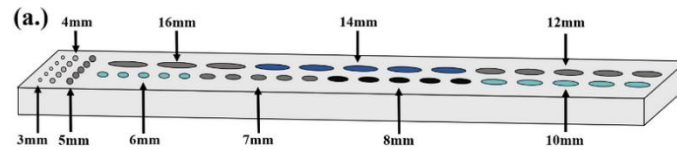
ORIGINAL RESEARCH

Quantitative Evaluation of PSMA PET Imaging using a Realistic Anthropomorphic Phantom and Shell-less Radioactive Epoxy Lesions

> Roberto Fedrigo, Dan J. Kadrmas, Patricia E. Edem, Lauren Fougner, Ivan S. Klyuzhin, M. Peter Petric, François Bénard, Arman Rahmim, Carlos Uribe

DOI: [10.21203/rs.3.rs-801202/v1](https://doi.org/10.21203/rs.3.rs-801202/v1) [Download PDF](#)

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Automatic Segmentation PSMA

Segmentation and analysis of PSMA-PET/CT images of prostate cancer

Ivan Klyuzhin, PhD



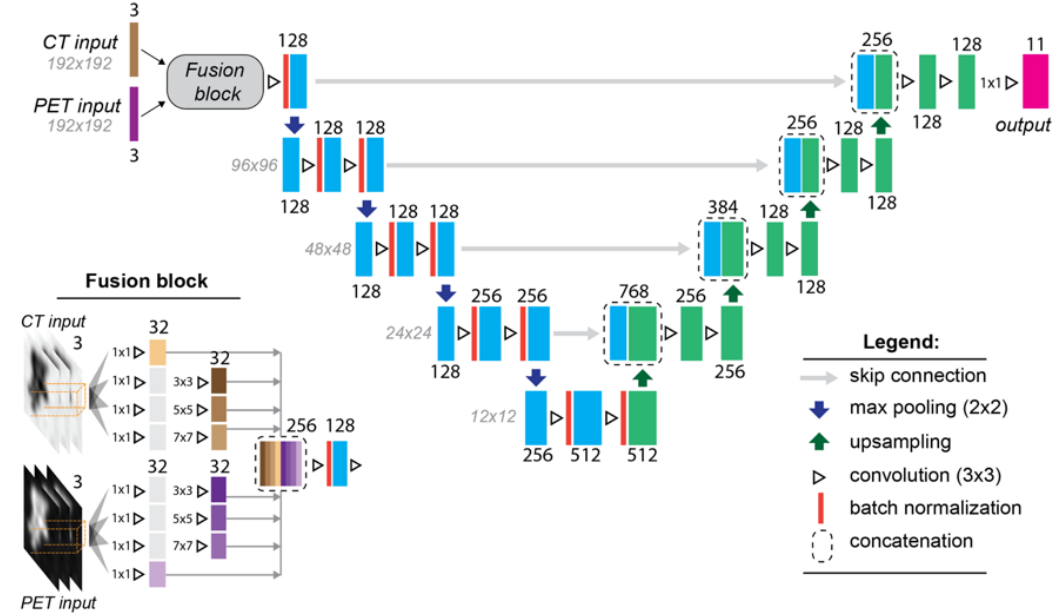
Measuring the tracer uptake in healthy organs can be used to optimize the drug dose and effectiveness on a personalized level.

[Violet et al., J Nucl Med., 2019]

- Manual segmentation of organs is very labor-intensive and subject to operator variability.

Our goal is to develop a fully-automated and robust method for healthy organ segmentation in PSMA PET/CT images.

Neural net architecture ("PSMA-Hornet")



Implementation:
Tensorflow/Keras

Activations:
leaky ReLU
sigmoid (output)

Total number of trainable parameters:
9,373,003

No dropout

Parameters per target:
937,300

Automatic Segmentation PSMA

Segmentation and analysis of PSMA-PET/CT images of prostate cancer

Ivan Klyuzhin, PhD



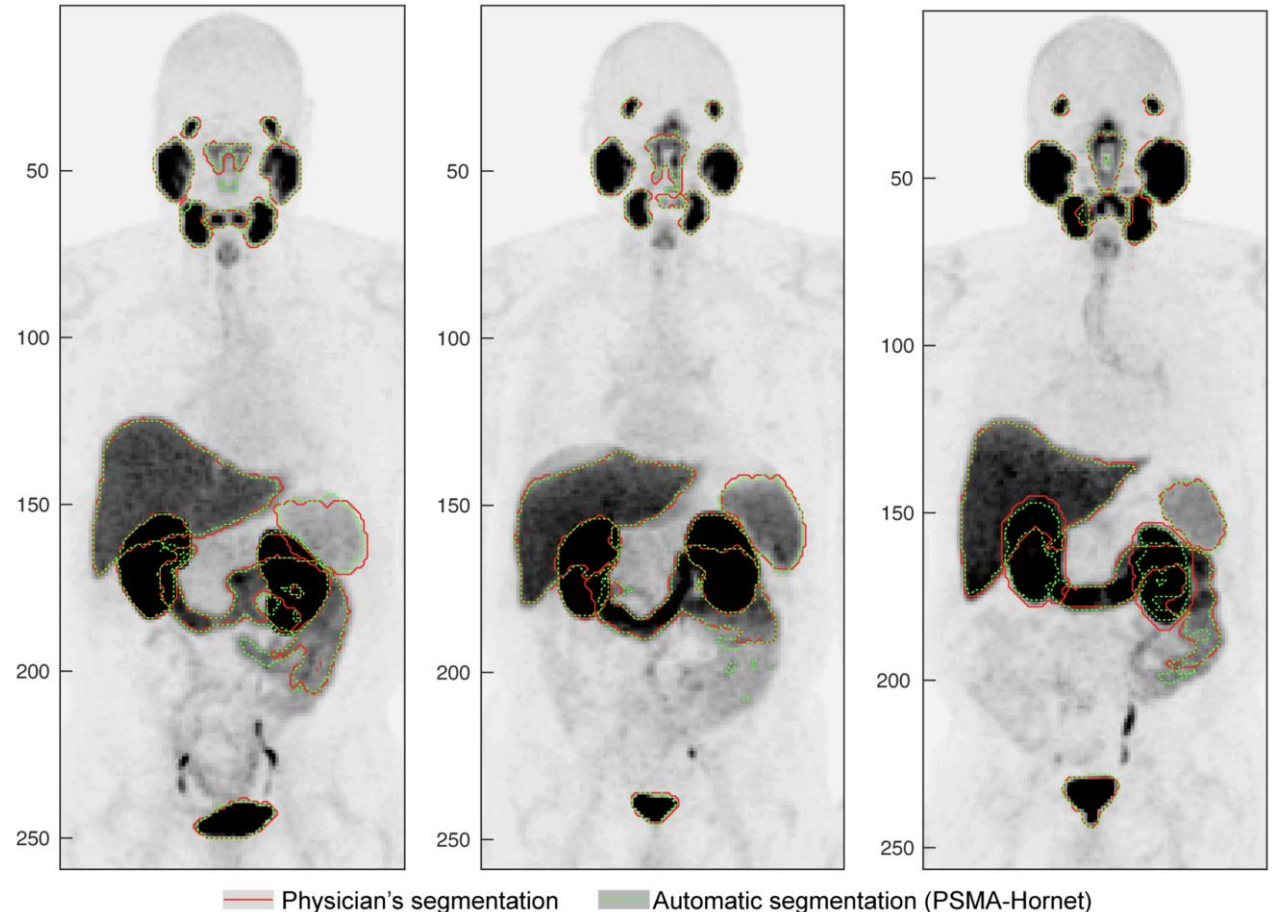
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PSMA-Hornet



Automatic Segmentation PSMA

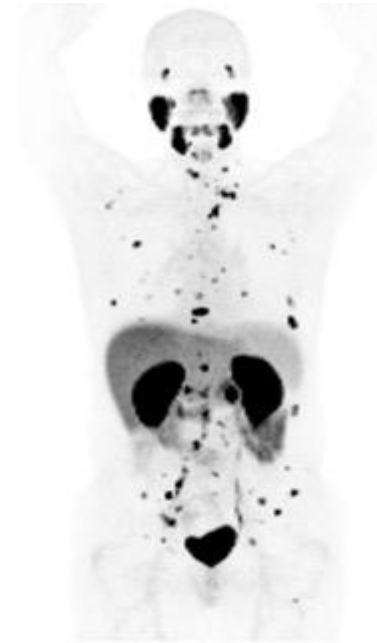
Segmentation and analysis of PSMA-PET/CT images of prostate cancer

Ivan Klyuzhin, PhD



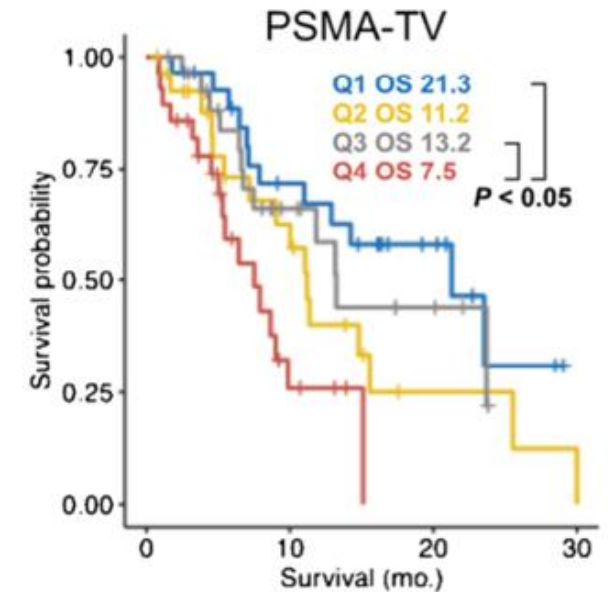
Total lesion volume (TV) was found to be significantly associated with prostate cancer survival – can be used to adjust therapy.

We are developing models and methods for fully-automated detection and segmentation of metastatic prostate cancer lesions.



PSMA Tracer: ^{18}F -DCFPyL

Schwarzenboeck et al., JNM, 2017



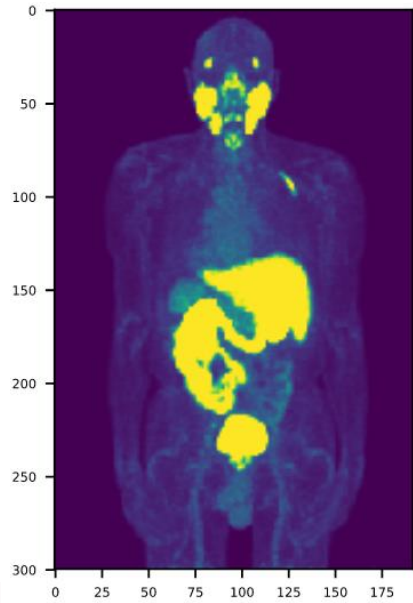
PSMA total tumor volume (TV) predicts survival.

Seifert et al., EJNMMI, Sep 2020

Automatic Segmentation PSMA

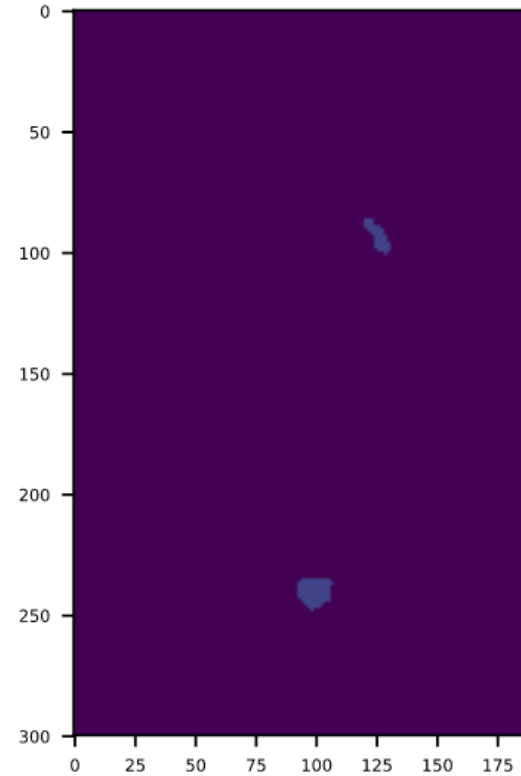
Segmentation and analysis of PSMA-PET/CT
images of prostate cancer

Ivan Klyuzhin, PhD

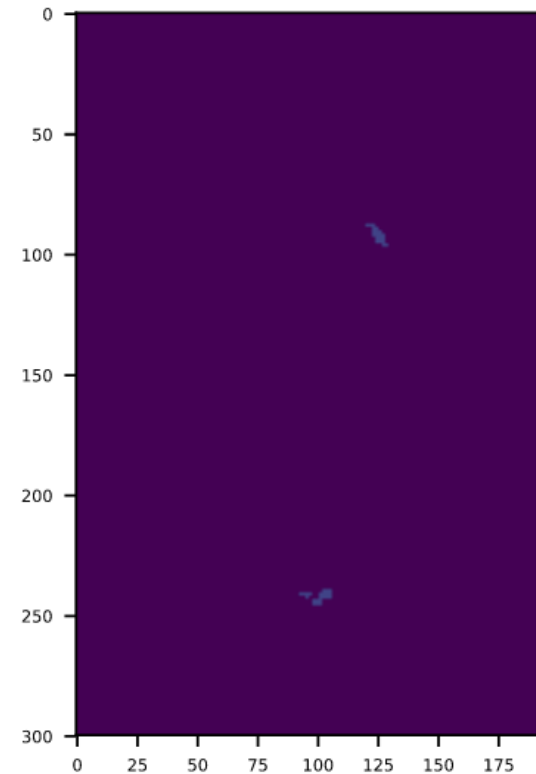


Lesion segmentation examples (U-net)

Ground truth



Prediction



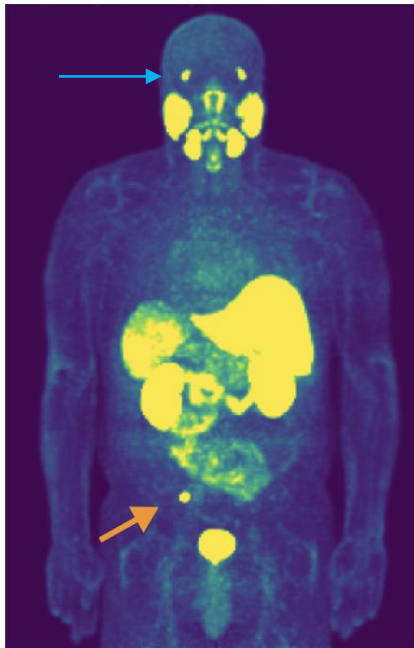
Automatic Segmentation PSMA

Segmentation and analysis of PSMA-PET/CT images of prostate cancer

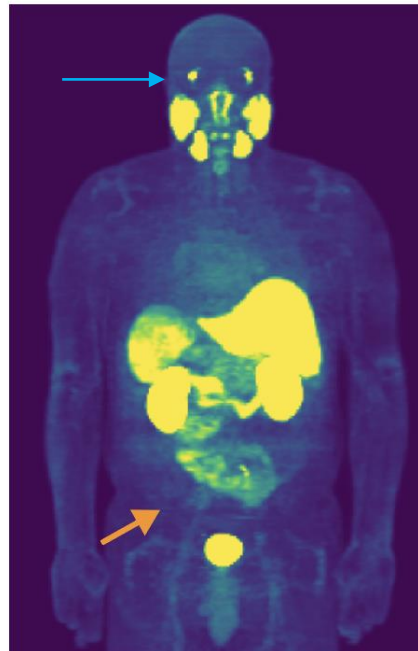
Ivan Klyuzhin, PhD



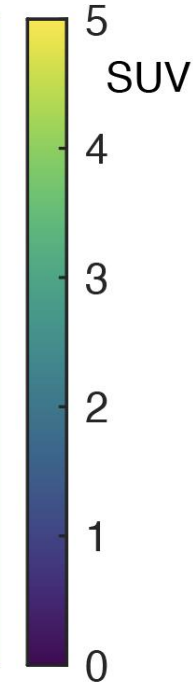
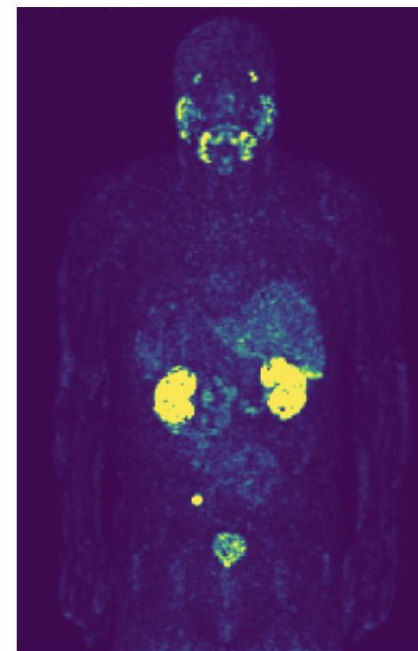
Original image



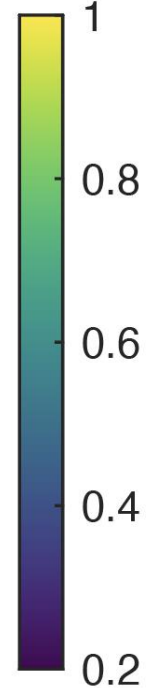
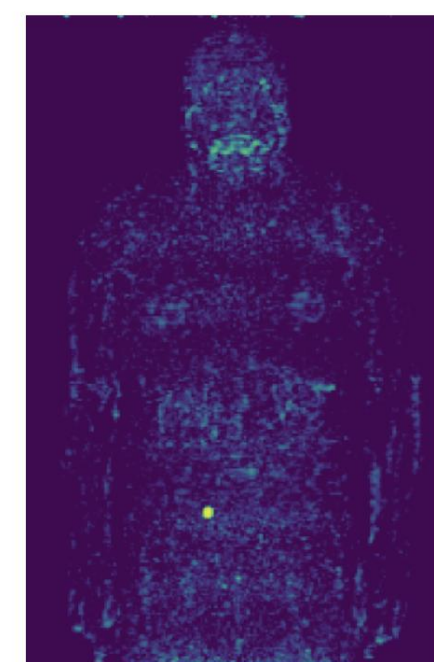
Predicted image



Residual image



Residual image
normalized



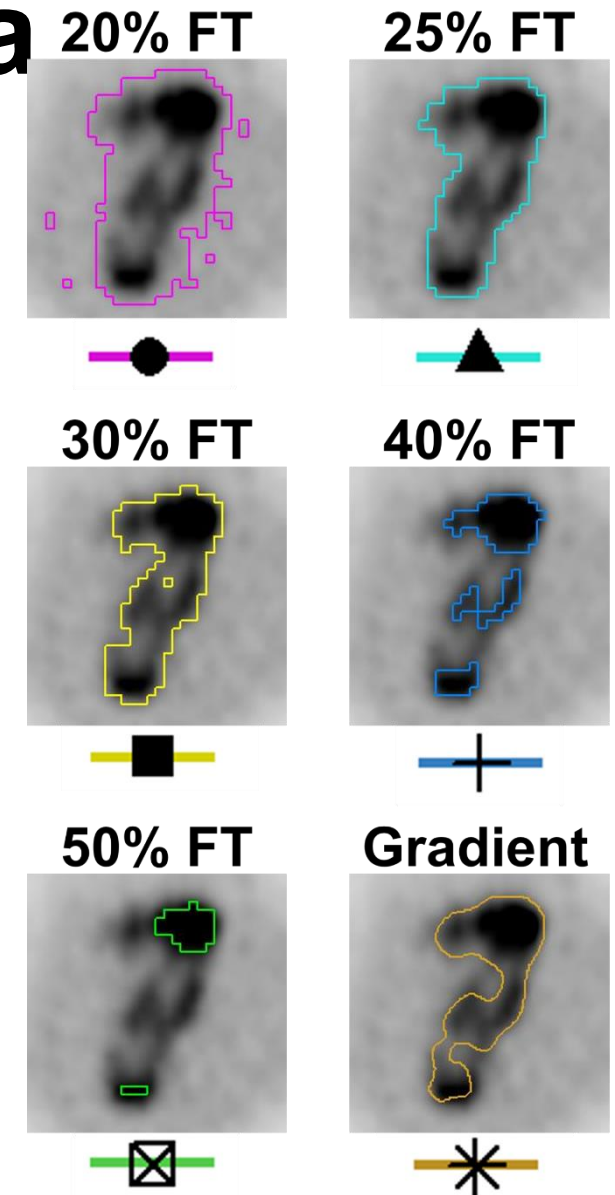
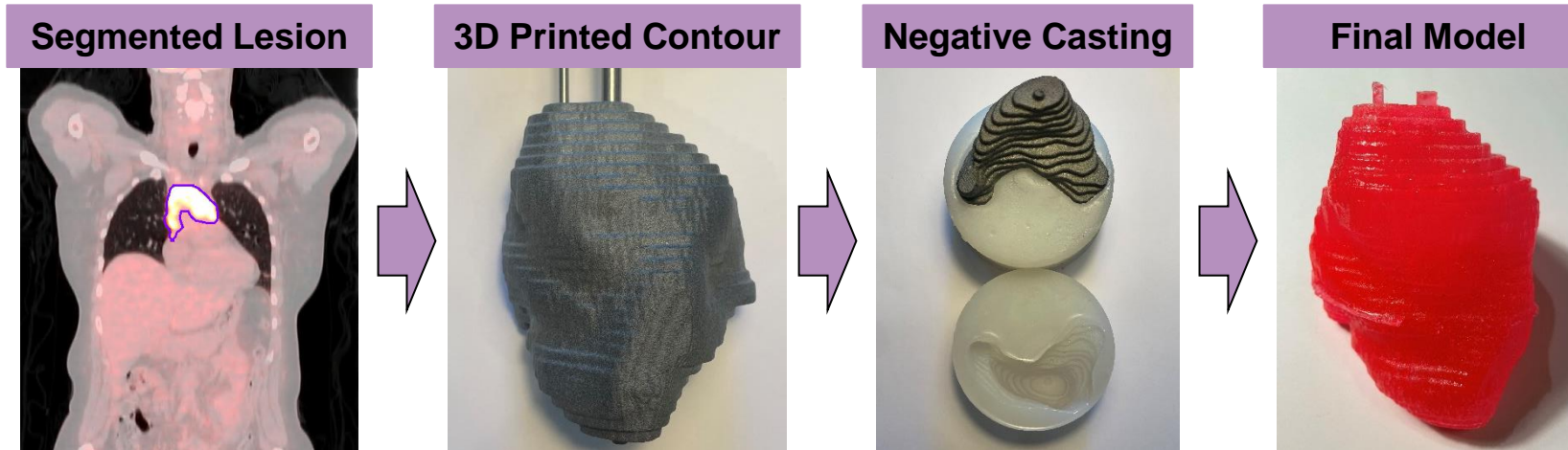
$$\frac{(\text{original} - \text{predicted})}{(\text{predicted})}$$

Quantitative Imaging Lymphoma

Negative-Cast Modelling for Oncology (NCMO)

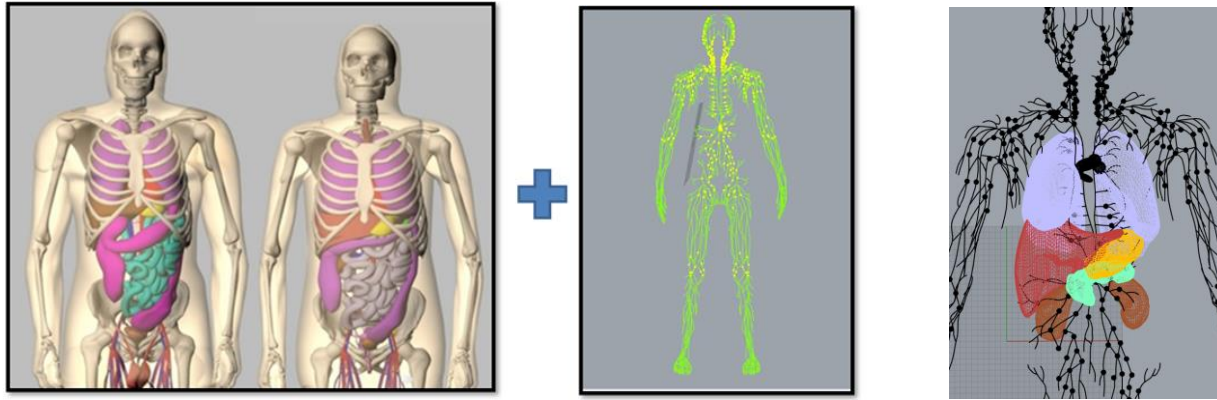
Realistic Tumour Modelling for Nuclear Medicine

- NCMO used to simulate bulky, heterogeneous tumours based on example cases from lymphoma patients



Roberto Fedrigo, Masters Student

Quantitative Imaging Lymphoma



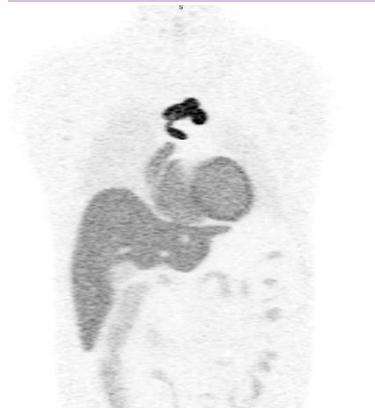
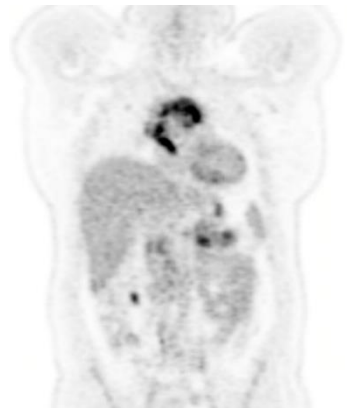
Development of the Lymphatic System in the 4D XCAT Phantom for Improved Multimodality Imaging Research

Roberto Fedrigo^{1,2}, Paul Segars³, Patrick Martineau⁴,
 Kerry J. Savage⁴, Carlos Uribe^{2,4}, Arman Rahmim^{1,2,4}
¹BC Cancer Research Institute, ²University of British Columbia,
³Duke University, ⁴BC Cancer



Patient

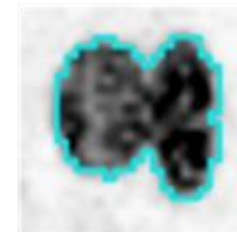
XCAT Simulation



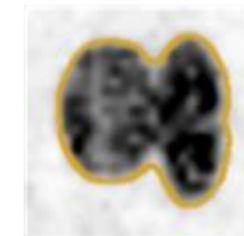
Best for Volume:

Best for Activity:

25% FT



Gradient



Roberto Fedrigo, Masters Student

Segmentation Lymphoma

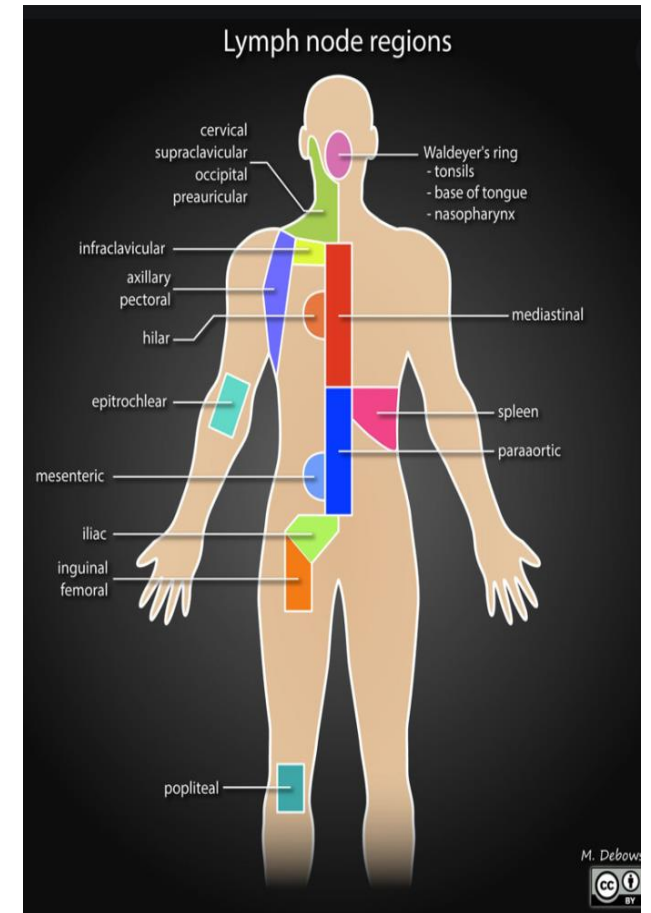
Fully automated segmentation of tumors
in PET and PET/CT images
using AI approaches

By: [Fereshteh Yousefirizi](#)



- Heterogeneous Presence:
 - lymphoma can be present in any of the over 500 lymph nodes
 - other lymphatic organs such as the bone marrow and spleen.
- Vast range of Treatments:
 - chemotherapy and radiation to newer immunotherapies

It is difficult to identify the patients who will not respond to therapy or the non responder patients



Segmentation Lymphoma

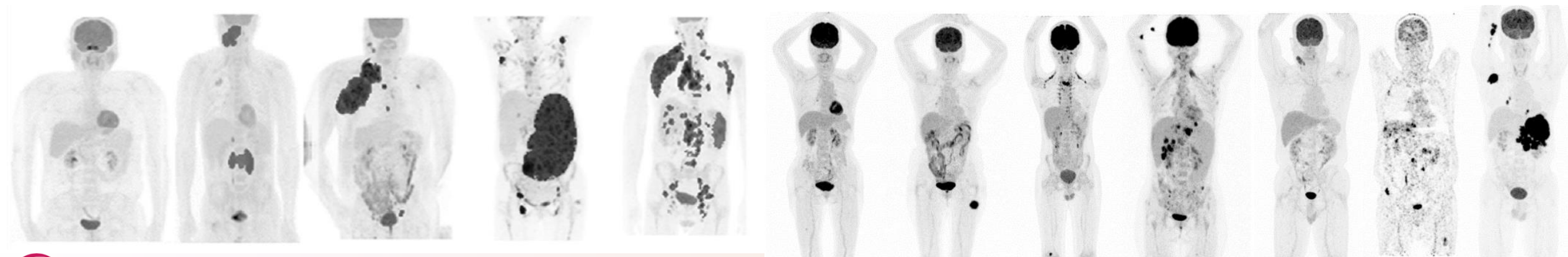
Fully automated segmentation of tumors
in PET and PET/CT images
using AI approaches

By: [Fereshteh Yousefirizi](#)



- Lymphoma lesion segmentation is a challenging task:

- 💡 Varied number, size, site and shape of lesions
- 💡 Heterogeneity and different degrees of glucose metabolism
- 💡 The high rate of false positive by normal organs (i.e. brain & bladder)



Segmentation Lymphoma

Fully automated segmentation of tumors
in PET and PET/CT images
using AI approaches

By: [Fereshteh Yousefirizi](#)

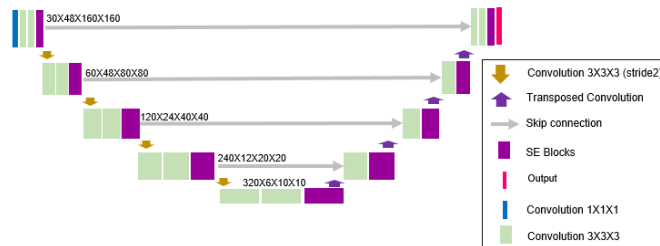


- Convolutional neural network with a hybrid loss function for fully automated segmentation of lymphoma lesions in FDG PET images, SPIE2022
- A cascaded deep network for automated tumor detection and segmentation in clinical PET imaging of diffuse large B-cell lymphoma, SPIE2022.
- Automated segmentation of diffuse large B cell lymphoma (DLBCL) lesions in [^{18}F]FDG-PET/CT images using Transfer learning, PILM2021

Segmentation Lymphoma

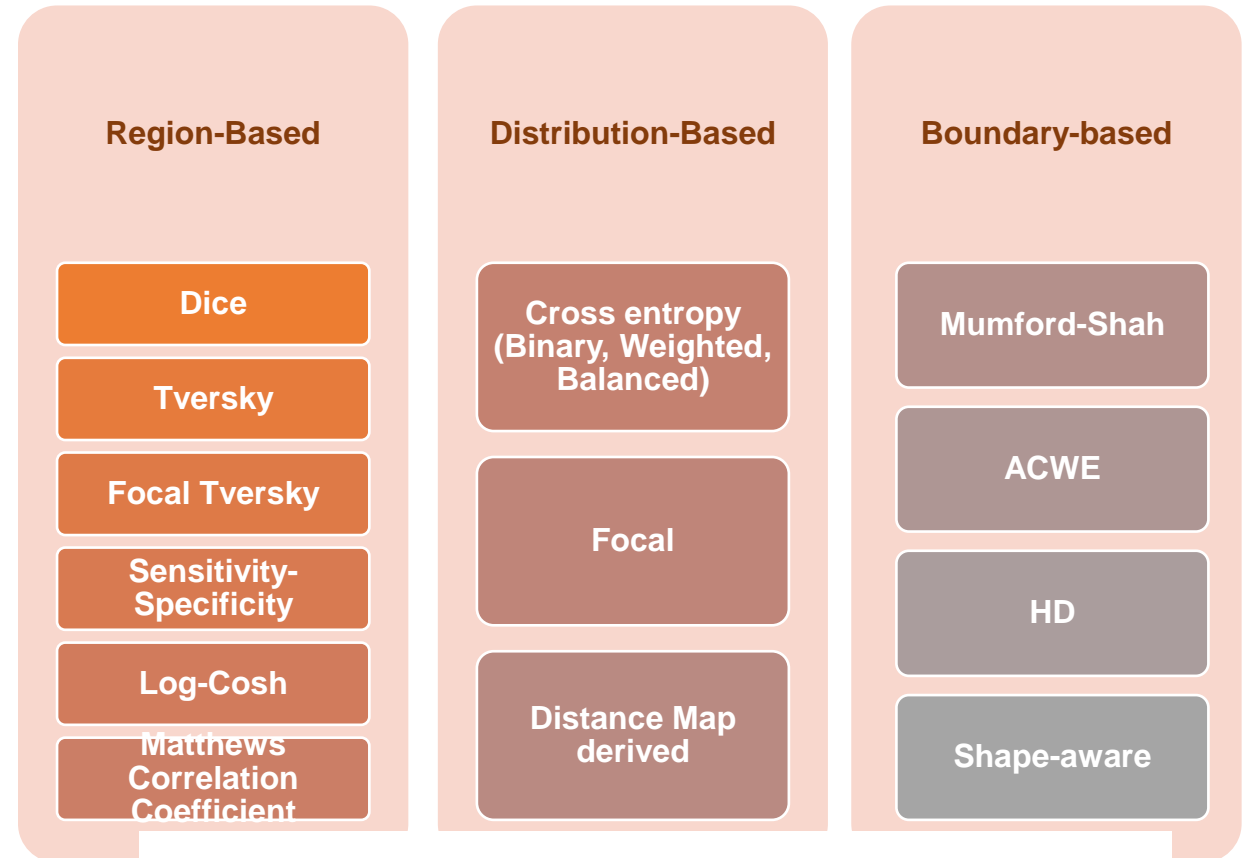
Fully automated segmentation of tumors
in PET and PET/CT images
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By: [Fereshteh Yousefirizi](#)



The **cross-entropy** loss and **Dice** loss or a combination of them are mainly used for semantic segmentation.

Three main **categories** of loss functions that have been used for **medical image segmentation**



$$Loss = L_{Distribution} + L_{Region} + L_{Boundray}$$

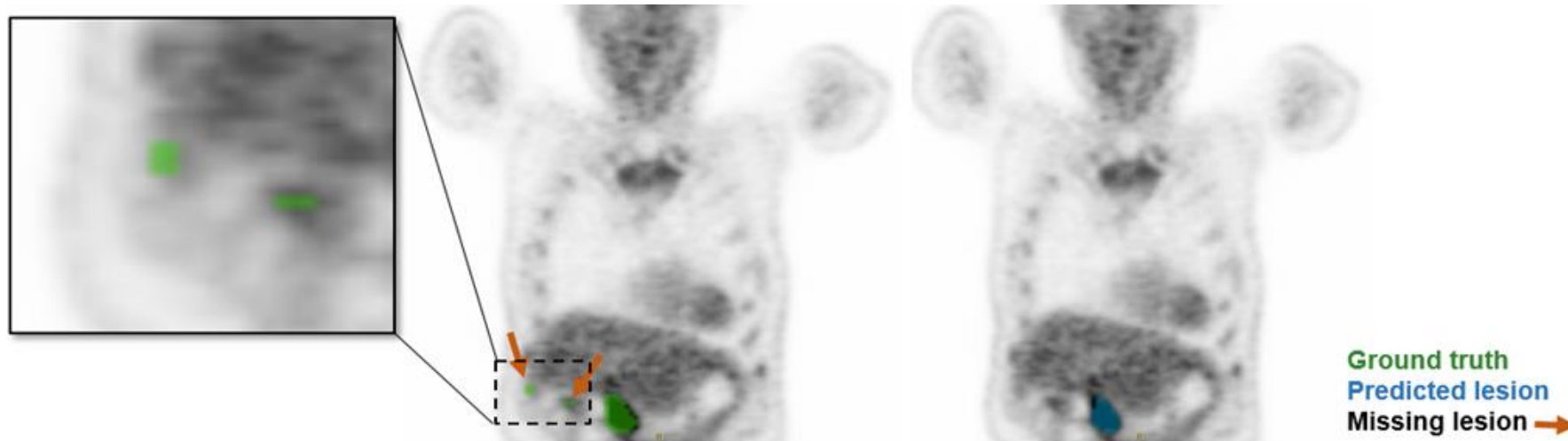
Segmentation Lymphoma

Fully automated segmentation of tumors
in PET and PET/CT images
using AI approaches

By: [Fereshteh Yousefirizi](#)



Loss function	Hyper-parameters	Dice	HD
Region	-	0.68 ± 0.21	24.6 ± 27.3
Distribution	$\alpha=0.25, \gamma=2$	0.72 ± 0.24	28.7 ± 19.2
Region + Distribution	$\alpha=0.25, \gamma=2$	0.75 ± 0.16	31.2 ± 21.9
Distribution + Region + Boundary	$\lambda=0.5, \delta=0.6, \gamma=0.5, \beta=10^{-7}$	0.77 ± 0.08	16.5 ± 12.5

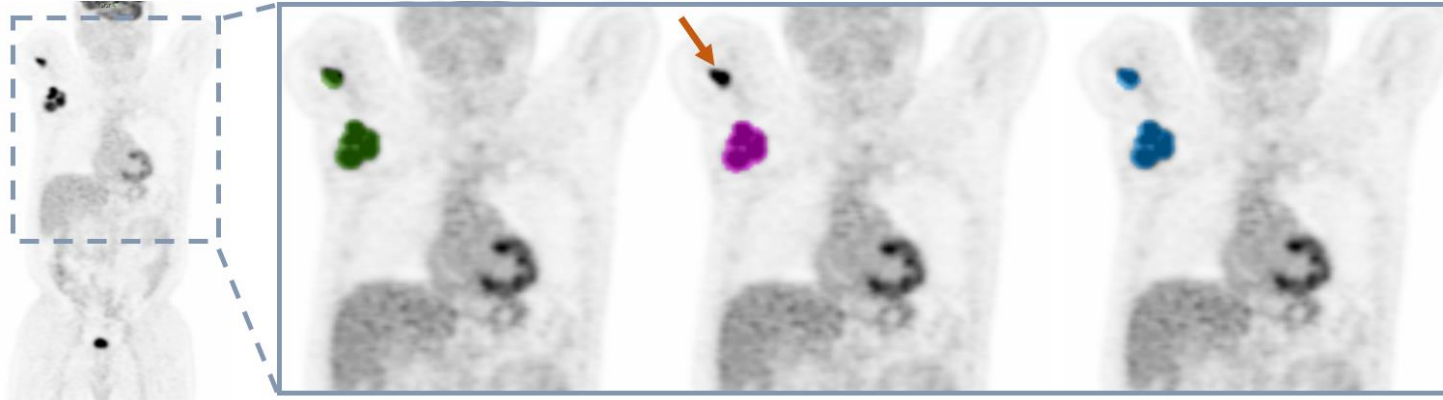


Segmentation Lymphoma

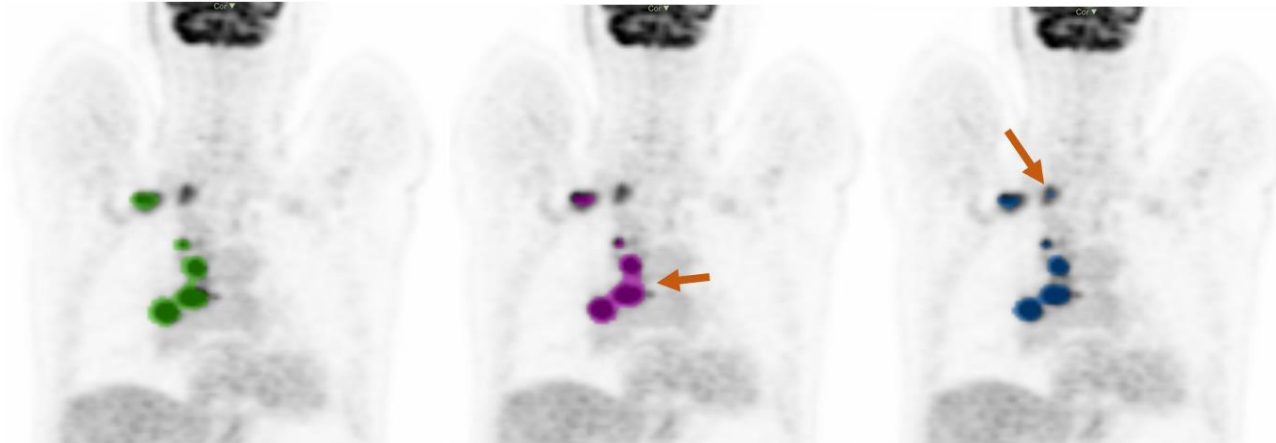
Fully automated segmentation of tumors
in PET and PET/CT images
using AI approaches

Approach1: Automated segmentation of diffuse large B cell lymphoma (DLBCL) lesions in [18F]FDG-PET/CT images using Transfer learning

By: [Fereshteh Yousefirizi](#)



(a) Original PET image (b) **ground truth mask** (c) trained on **PMBCL data** (d) trained on **H&N data**



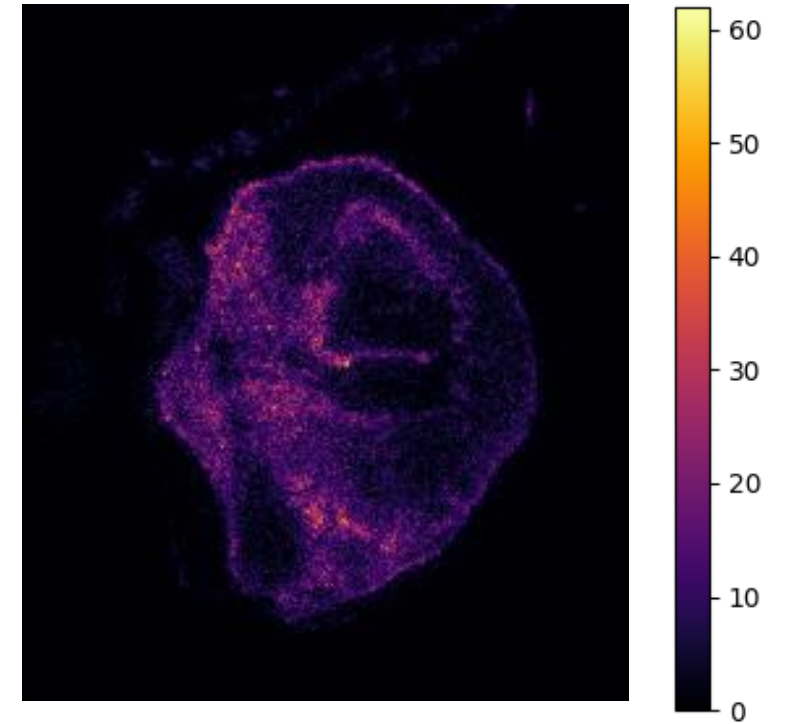
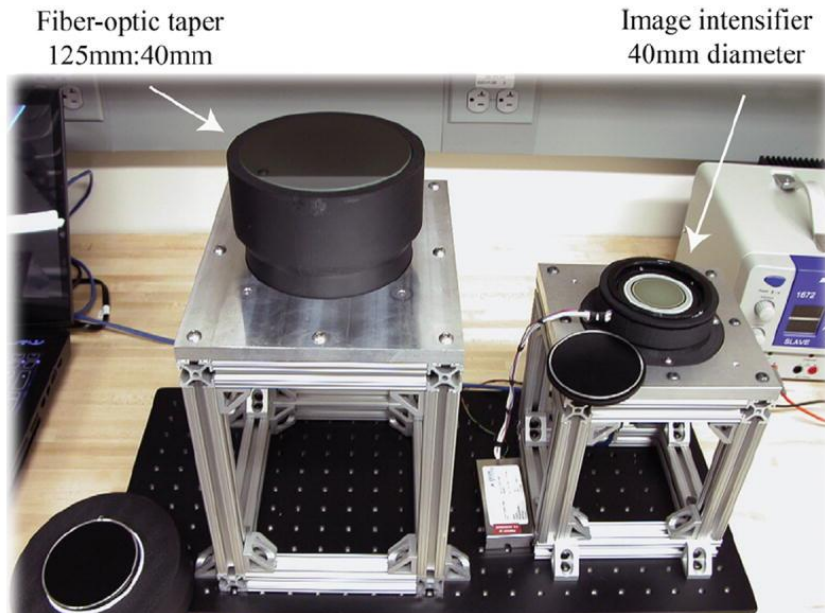
(a) **ground truth mask** (b) trained on **PMBCL data** (c) trained on **H&N data**

Model	DSC (mean ± std)
trained on H&N	0.71±0.14
trained on PMBCL	0.54±0.08

Multiscale Dosimetry

Quantitative Ex Vivo Imaging of ^{225}Ac with the iQID Alpha Camera

Cassandra Miller^{1,2}, Julie Rousseau³, Jason Crawford⁴, Brian Miller^{5,6}, François Bénard^{3,8,9}, Arman Rahmim^{1,2,8}, Carlos Uribe^{7,9}



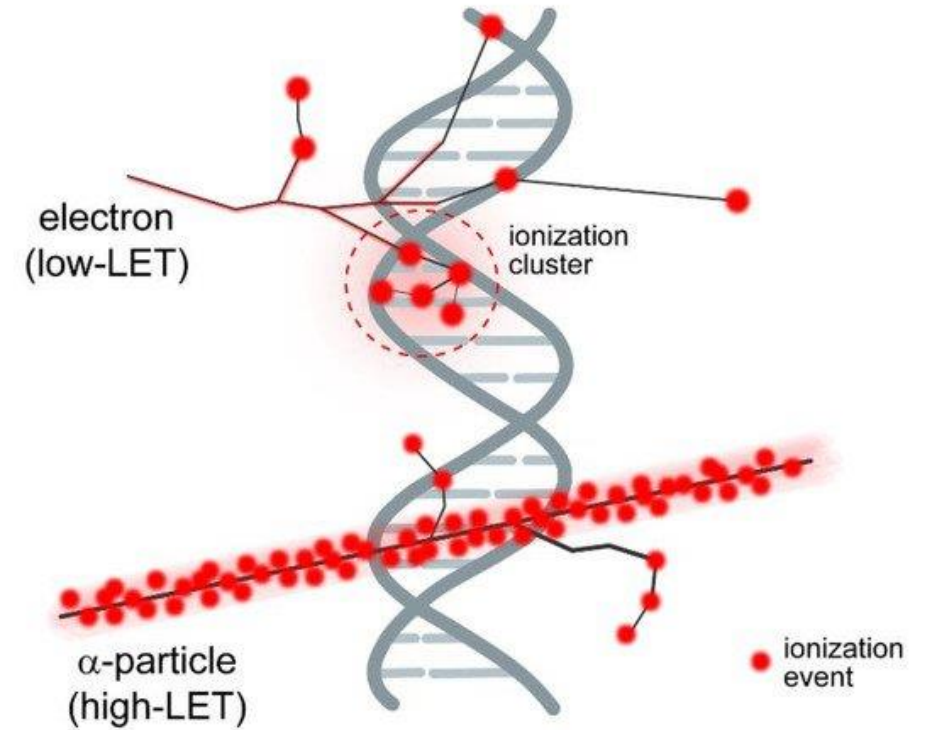
Intratumoural distribution of an ^{225}Ac labelled antibody in mouse tissue section

Multiscale Dosimetry

The Impact of Cell Shape on the Doses Delivered to the Nucleus from ^{177}Lu -labelled Radiotracers

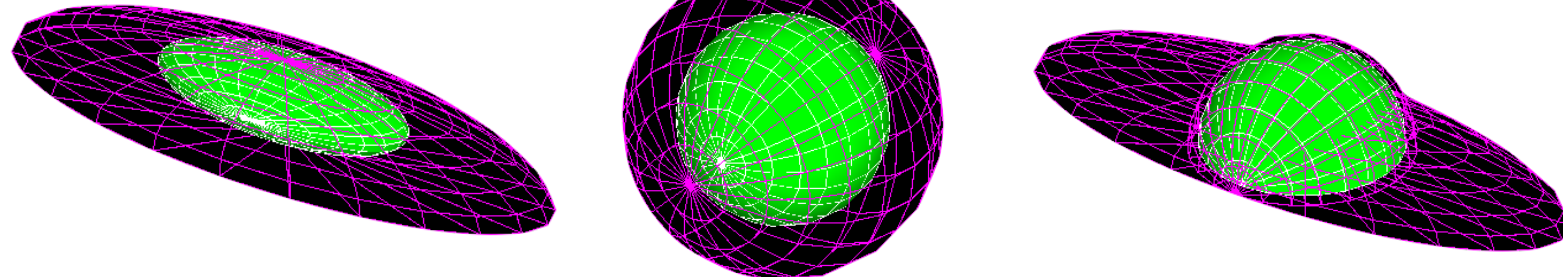
Cassandra Miller^{1,2}, Guillaume Chaussé³, Julia Brosch-Lenz^{1,4}, François Bénard^{3,5,6}, Arman Rahmim^{1,2,6}, and Carlos Uribe^{3,6}

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Macroscopic dosimetry **omits** the physics & biology that occurs on the **microscopic** and **cellular** level

There are microscopic variations in absorbed doses between radioisotopes and pharmaceuticals which may not be seen at the organ-level

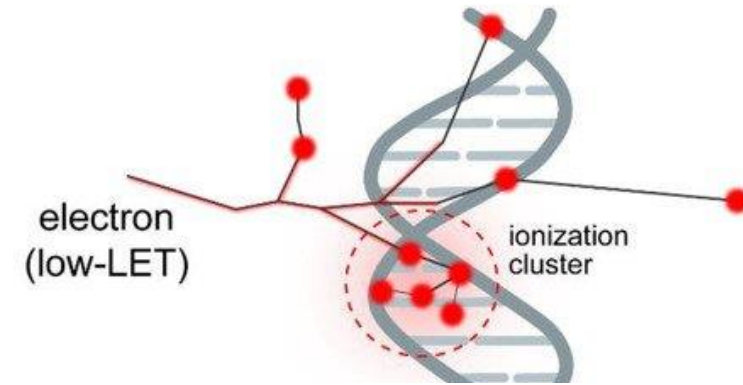


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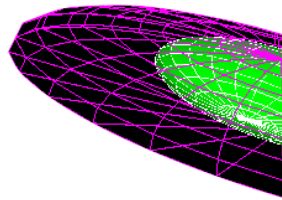
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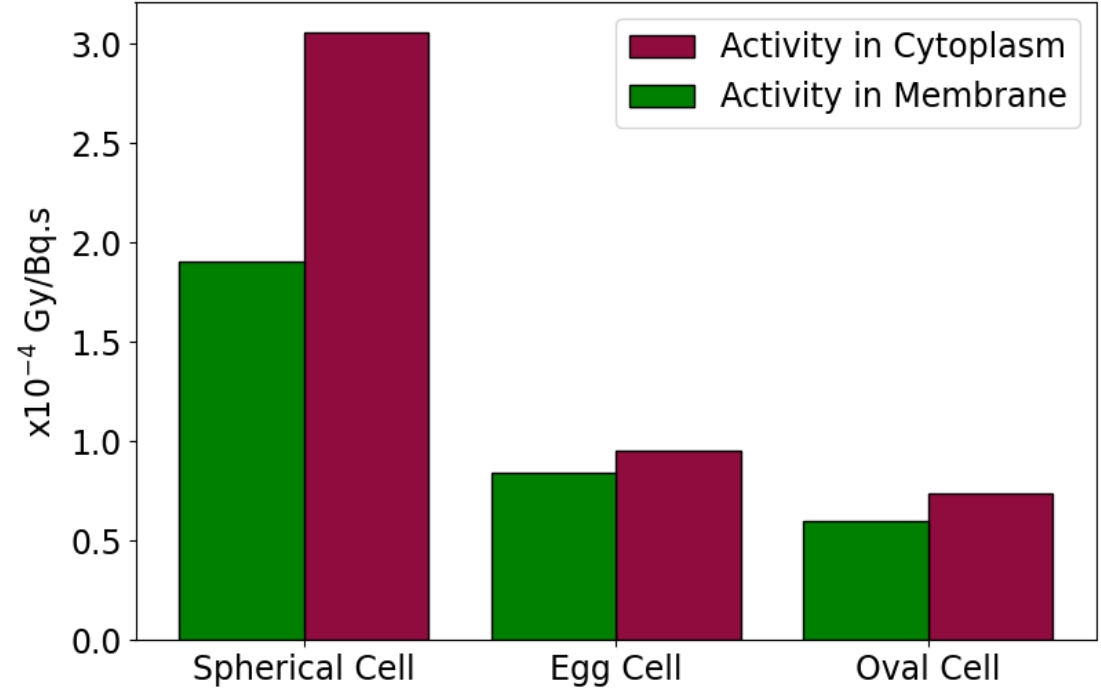


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S-Value to Nucleus

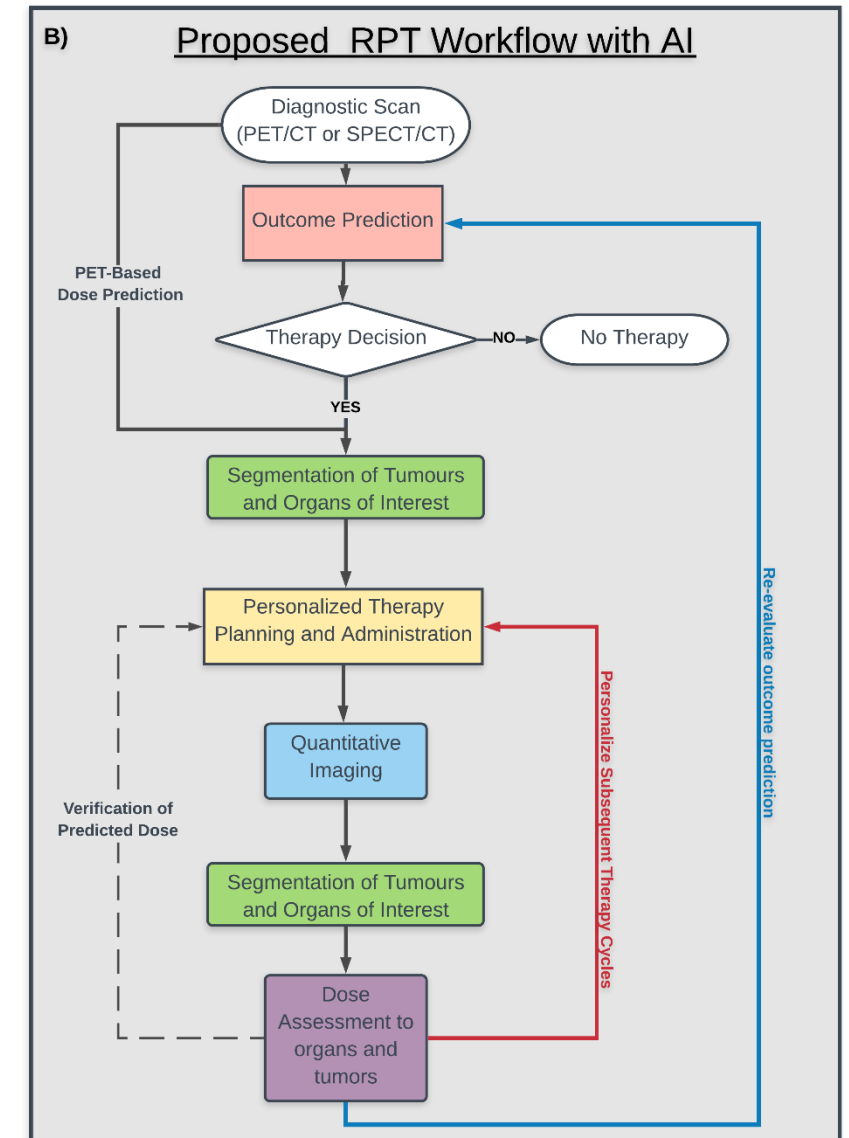


Conclusions and Acknowledgements

Conclusions



- Nuclear medicine detectors (i.e. PET and SPECT) allow us to gather information of
- Quantitative imaging is needed for accurate dosimetry and outcome prediction
- AI is a powerful tool that can help us with the different steps of the dosimetry workflow to make personalized dose assessments routine in clinical practice



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SOCIETY OF
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Thank you!



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