

LATEST DEVELOPMENTS IN CZT SPECT IMAGING

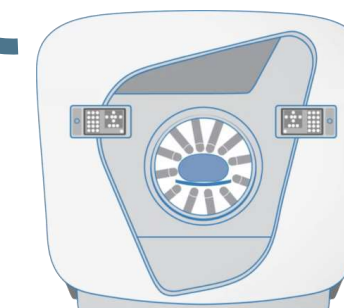
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Director of Nuclear Medicine Physics, Brigham and Women's Hospital
Faculty in Radiology, Harvard Medical School*



DISCLOSURE

I have no relevant financial interests to disclose

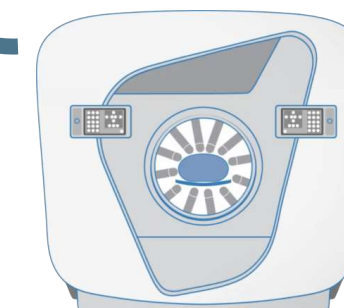




LEARNING OBJECTIVES

At the end of this presentation, you will be able to,

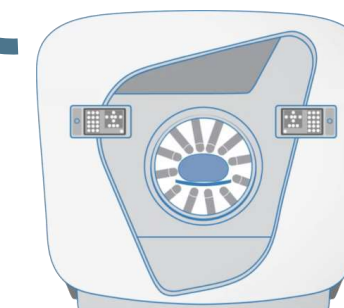
- A) Recognize the specificity of CZT-based SPECT systems**
- B) Identify the advantages of CZT detector technology for imaging**
- C) Categorize the clinical applications of CZT-based SPECT and SPECT/CT systems**





OUTLINE

- I. CZT DETECTORS FOR SPECT
- II. CARDIAC DEDICATED CZT SPECT
- III. FULL-RING CZT SPECT/CT
- IV. CLINICAL APPLICATIONS AND ADDED VALUE
- V. ABSOLUTE QUANTITATION CALIBRATION
- VI. IMPACT OF THE **CALIBRATION ACCURACY** ON QUANTITATION





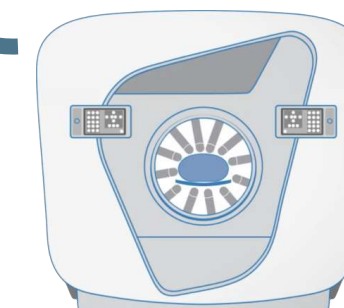
Cadmium Zinc Telluride (CZT)

Characteristics

- Semiconductor at room temperature
- Direct Conversion** of the γ -ray energy into an electronic signal
- Higher Density** than conventional NaI(Tl): **5.8** Vs **3.7** g/cm³. However, due to cost considerations, CZT detectors are thinner (**5 to 7.3 mm**) than conventional (**9.5 mm**).
Stopping Power is very similar (~95% for ^{99m}Tc) [1,2].

[1] Wells, *et al.* J. Nucl. Cardiol. 2020

[2] Slomka, *et al.* J. Nucl. Med., 2019





Cadmium Zinc Telluride (CZT)

Characteristics

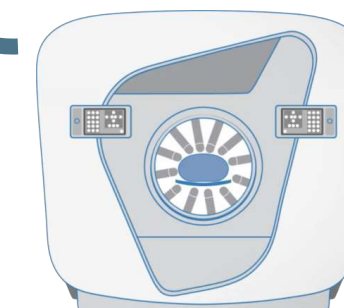
- ❑ *Semi-Conductor*
- ❑ *Direct Conversion of the γ -ray energy into an electronic signal*
- ❑ *Stopping Power is very similar to NaI(Tl)*

Performance compared to conventional NaI(Tl)+PMT detector

- ❑ **Better Energy Resolution: 5-6% Vs 9-10% for ^{99m}Tc [1-3]**

→ Improves *scatter discrimination* with narrower energy window

[1] Wells, *et al.* J. Nucl. Cardiol. 2020
[2] Slomka, *et al.* J. Nucl. Med., 2019
[3] Hutton, *et al.* Clin. Transl. Imaging, 2018





Cadmium Zinc Telluride (CZT)

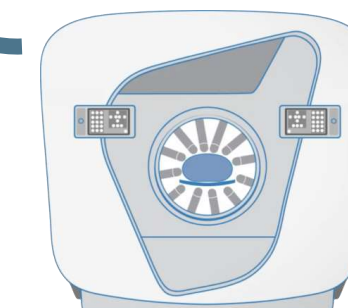
Characteristics

- Semi-Conductor*
- Direct Conversion of the γ -ray energy into an electronic signal*
- Stopping Power is very similar to NaI(Tl)*

Performance compared to conventional NaI(Tl)+PMT detector

- Better Energy Resolution
- Improved **Detector Spatial Resolution** (= pixel size **~2.46 mm**, typically) Vs **~3.8 mm** for ^{99m}Tc [1]

[1] Hutton, et al. Clin. Transl. Imaging, 2018





Cadmium Zinc Telluride (CZT)

Characteristics

- ❑ *Semi-Conductor*
- ❑ *Direct Conversion of the γ -ray energy into an electronic signal*
- ❑ *Stopping Power is very similar to NaI(Tl)*

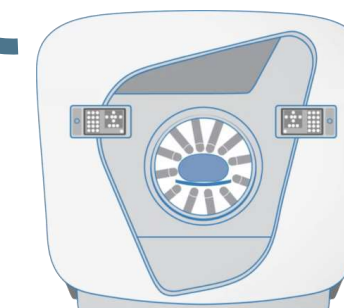
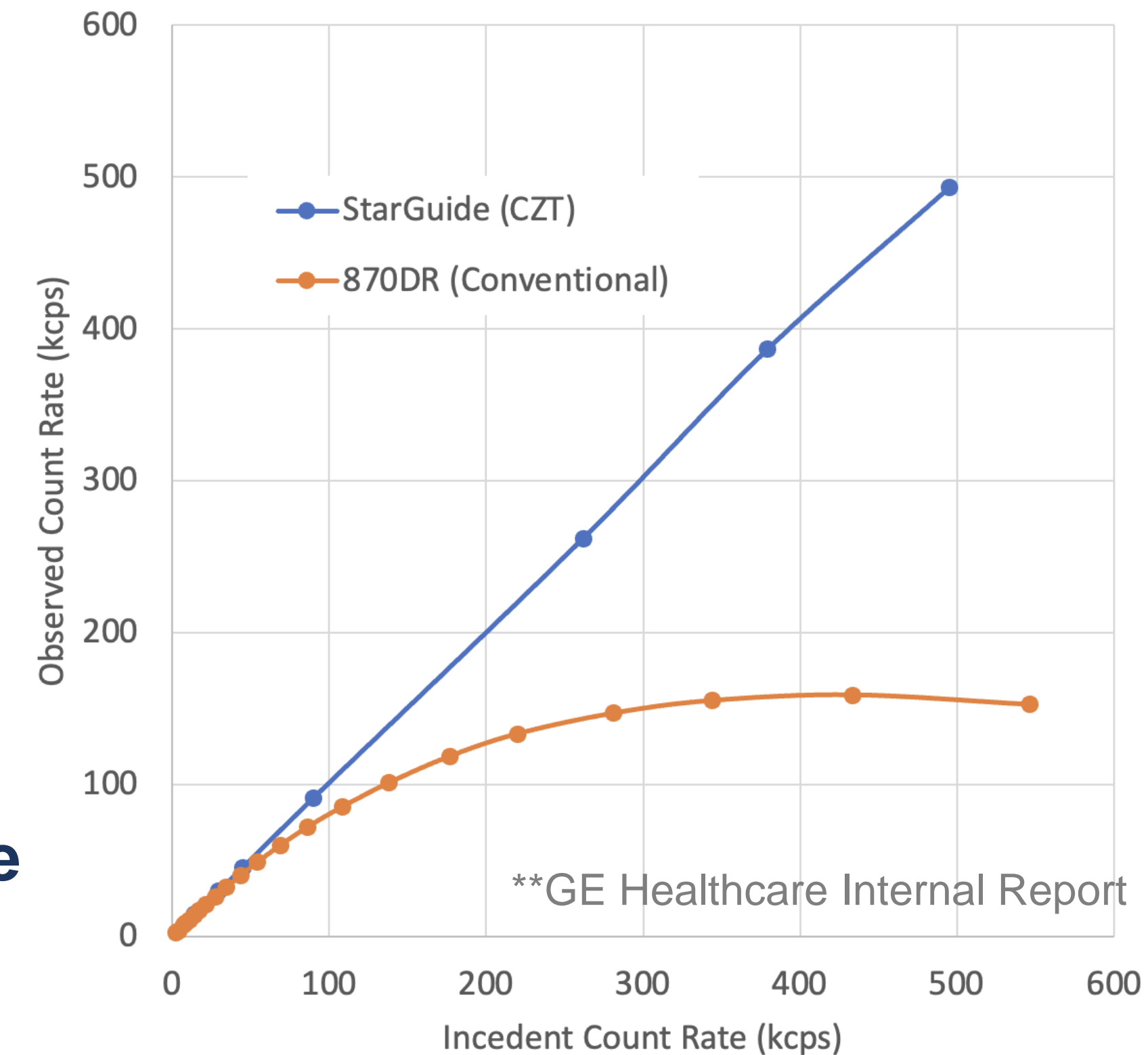
Performance compared to conventional NaI(Tl)+PMT detector

- ❑ *Better Energy Resolution*
- ❑ *Improved Detector Spatial Resolution*
- ❑ **High Count-Rate Capability**

➔ **No dead time or detector saturation in the clinical range**
 < 1% count rate loss Vs 20% loss \geq 250 kcps [1]

[1] GE Healthcare. NM870 SPECT/CT specification sheet

StarGuide Count Rate Performance



Cadmium Zinc Telluride (CZT)

Characteristics

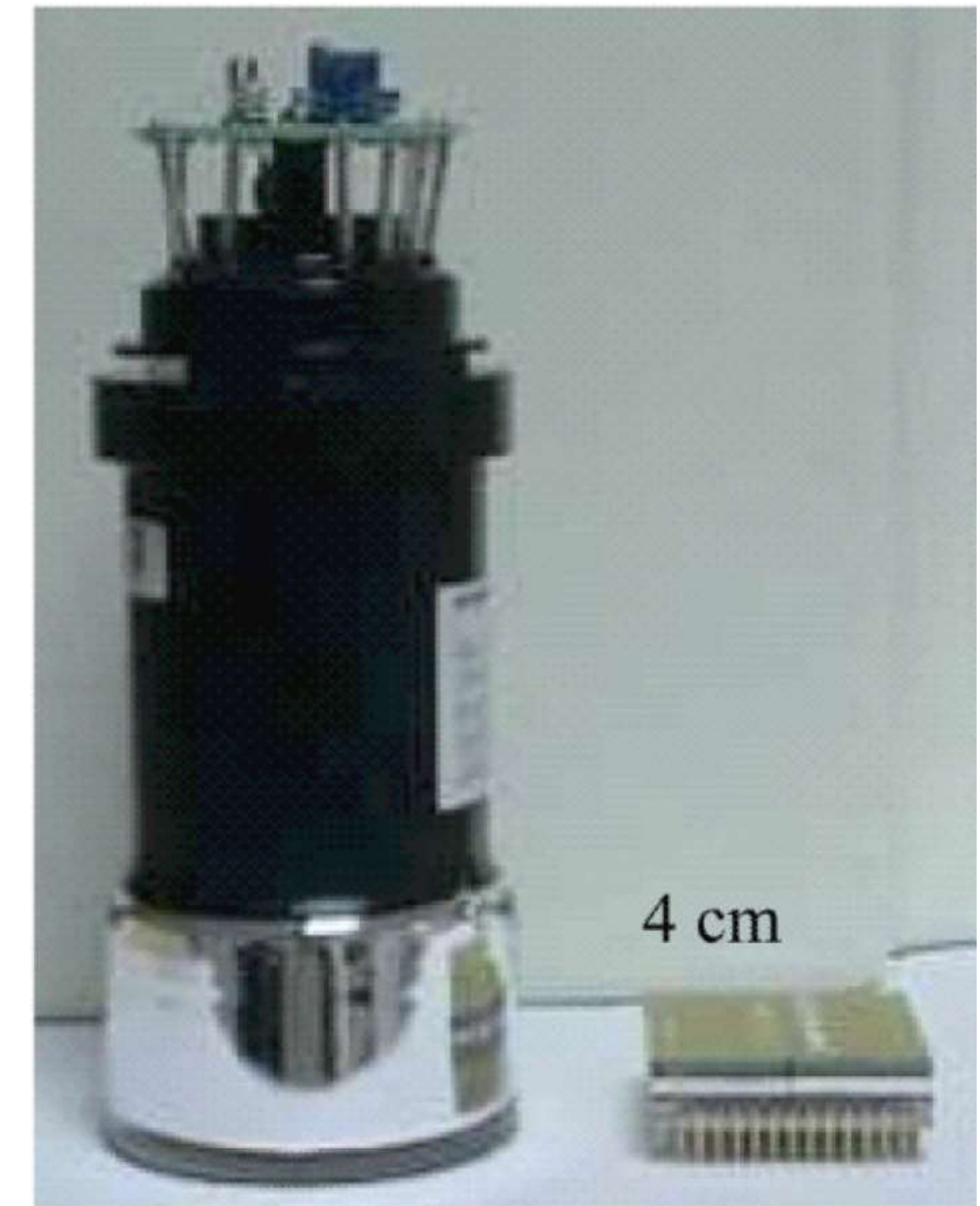
- Semi-Conductor*
- Direct Conversion of the γ -ray energy into an electronic signal*
- Stopping Power is very similar to NaI(Tl)*

Performance compared to conventional NaI(Tl)+PMT detector

- Better Energy Resolution**
- Improved Detector Spatial Resolution**
- Enhanced Count-Rate Capability**
- Higher Cost**
- Compact Detector** facilitates more **efficient SPECT System Design**

NaI(Tl)
+ PMT

CZT



Adapted from Peterson and Furenlid.
Phys. Med. Biol., 2011

MYOSPECT® (GENERAL ELECTRIC HEALTHCARE)

- ❑ 19 or 9 (*MyoSPECT ES*) CZT-based detectors
- ❑ Multi-Pinhole Collimation
- ❑ Each detector focuses on the cardiac region

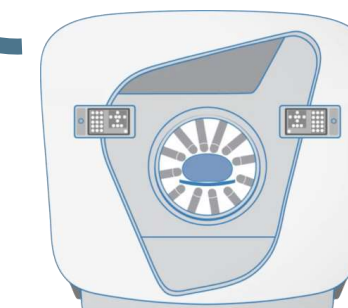


D-SPECT® (SPECTRUM DYNAMICS MEDICAL)

- ❑ 9 (*Cardio*) or 6 (*Vista*) CZT-based detectors
- ❑ Focusing Parallel-Hole Collimation
- ❑ Each detector focuses on the cardiac region



Courtesy of Spectrum Dynamics Medical





FULL-RING CZT SPECT/CT

NOVEL SYSTEM DESIGNS

- ❑ 360-degree acquisition
- ❑ Closer patient contouring compared to conventional systems
- ❑ Adaptive Imaging for a broad range of clinical studies

STARGUIDE® (GENERAL ELECTRIC)

- ❑ 12 CZT-based detectors
7.25 mm thick; Up to 270 keV
- ❑ Swiveling Parallel-Hole Collimation
- ❑ Optical Scan Guided



VERITON-CT® (SPECTRUM DYNAMICS)

- ❑ 12 CZT-based detectors
200 Series: 6.0 mm thick; Up to 200 keV
400 Series: 7.3 mm thick; Up to 400 keV
- ❑ Swiveling Parallel-Hole Collimation
- ❑ Detectors can focus on a VOI Region

First US Install!





FULL-RING CZT SPECT/CT

VERITON-CT[®]

SPECT

- ❑ 12 CZT Swiveling Detectors
- ❑ 360-degree Acquisition
- ❑ Adaptive Imaging
- ❑ Parallel-Hole Collimators

CT

- ❑ 16, 64, 128 slices (0.625 mm)
- ❑ Low-dose CT for SPECT AC
- ❑ Diagnostic AC

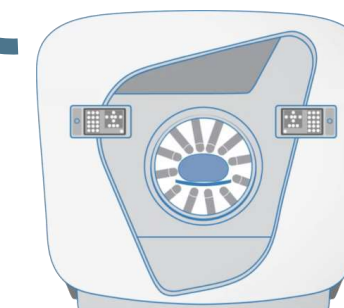
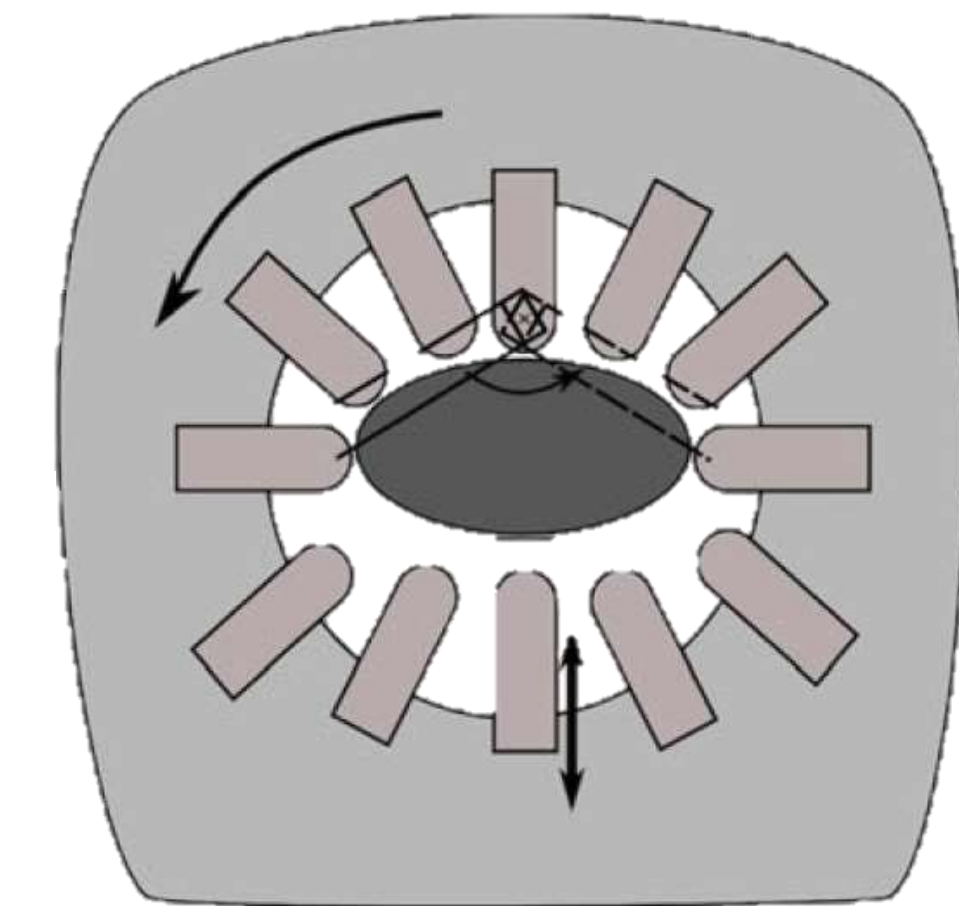
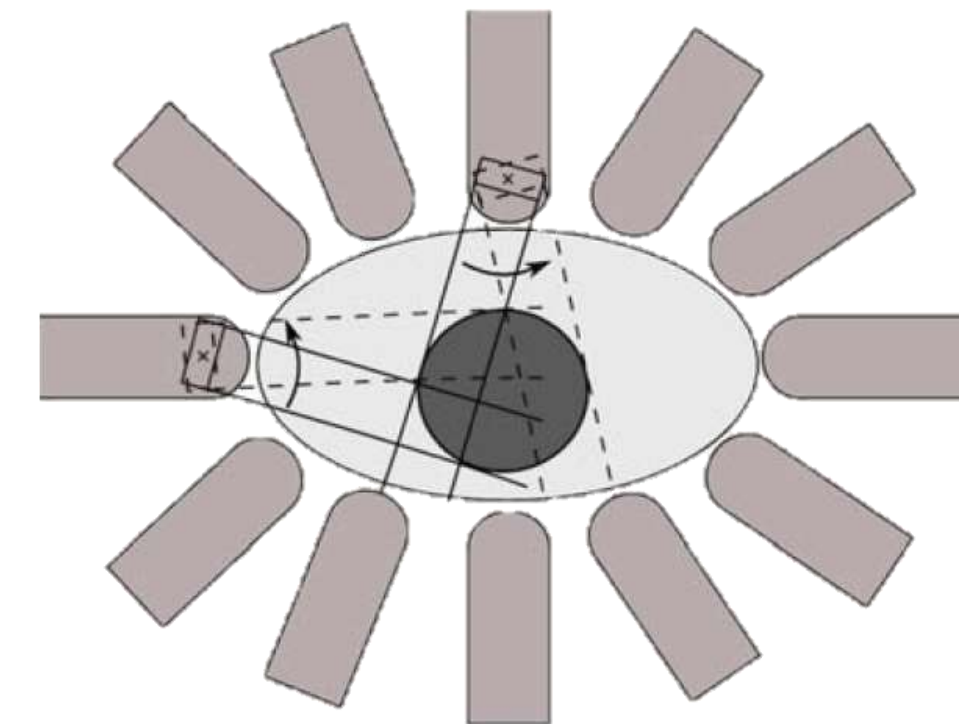


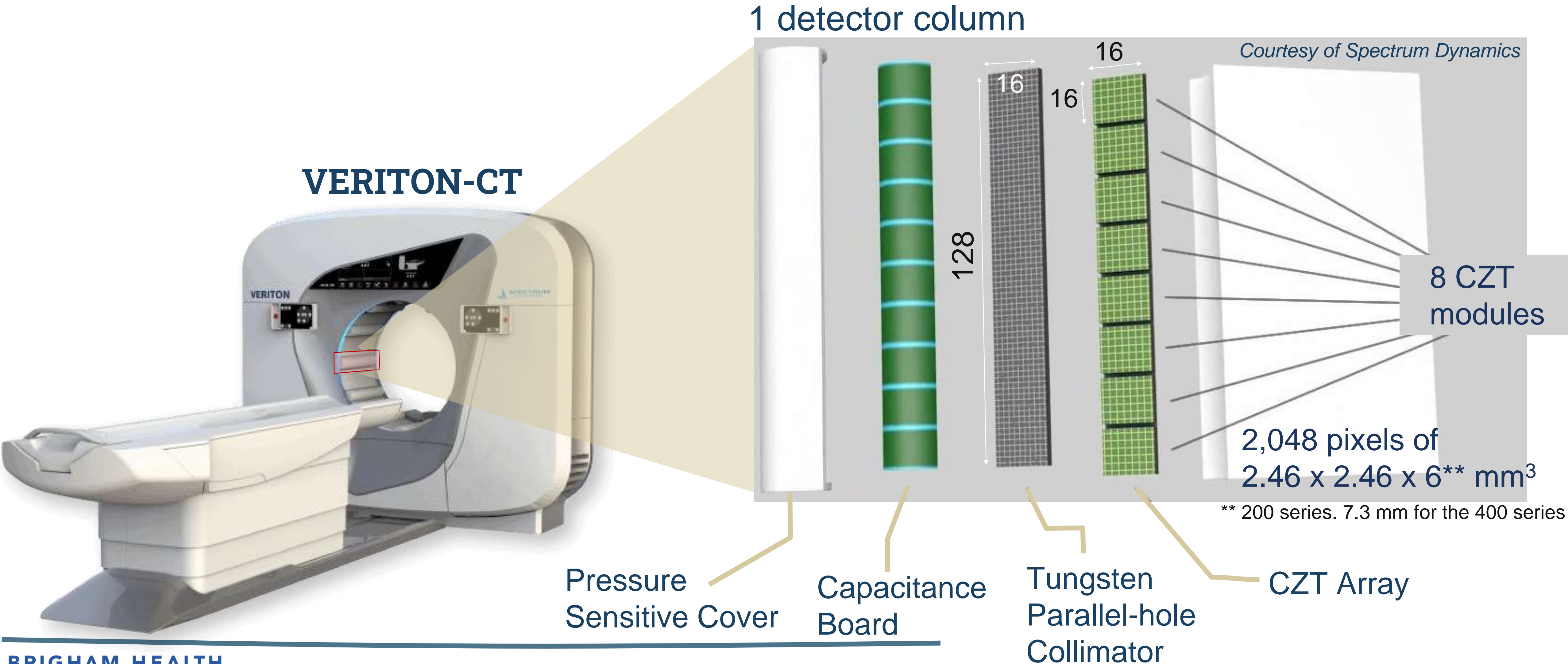
Broad Range of Clinical Applications

Focused Scan

3D Dynamic Scan

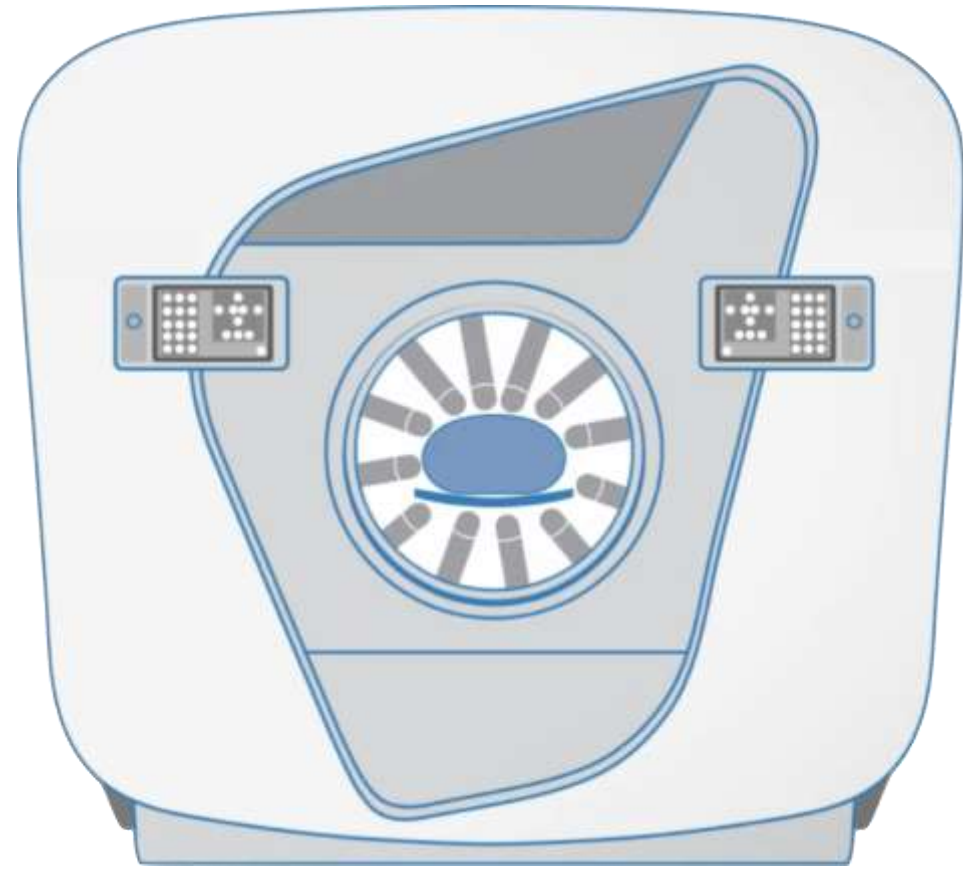
Whole-Body Scan





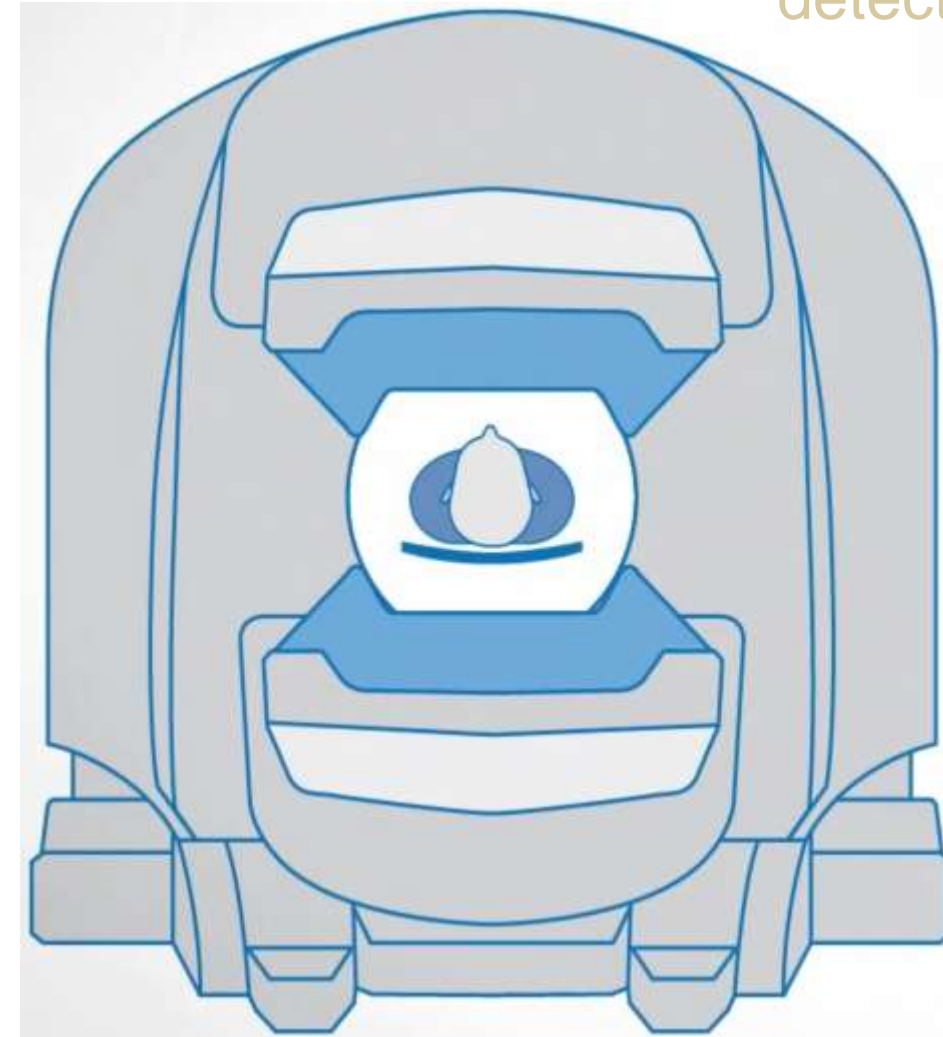
Improved Imaging Performance

High Sensitivity and Improved Spatial Resolution compared to Conventional Systems

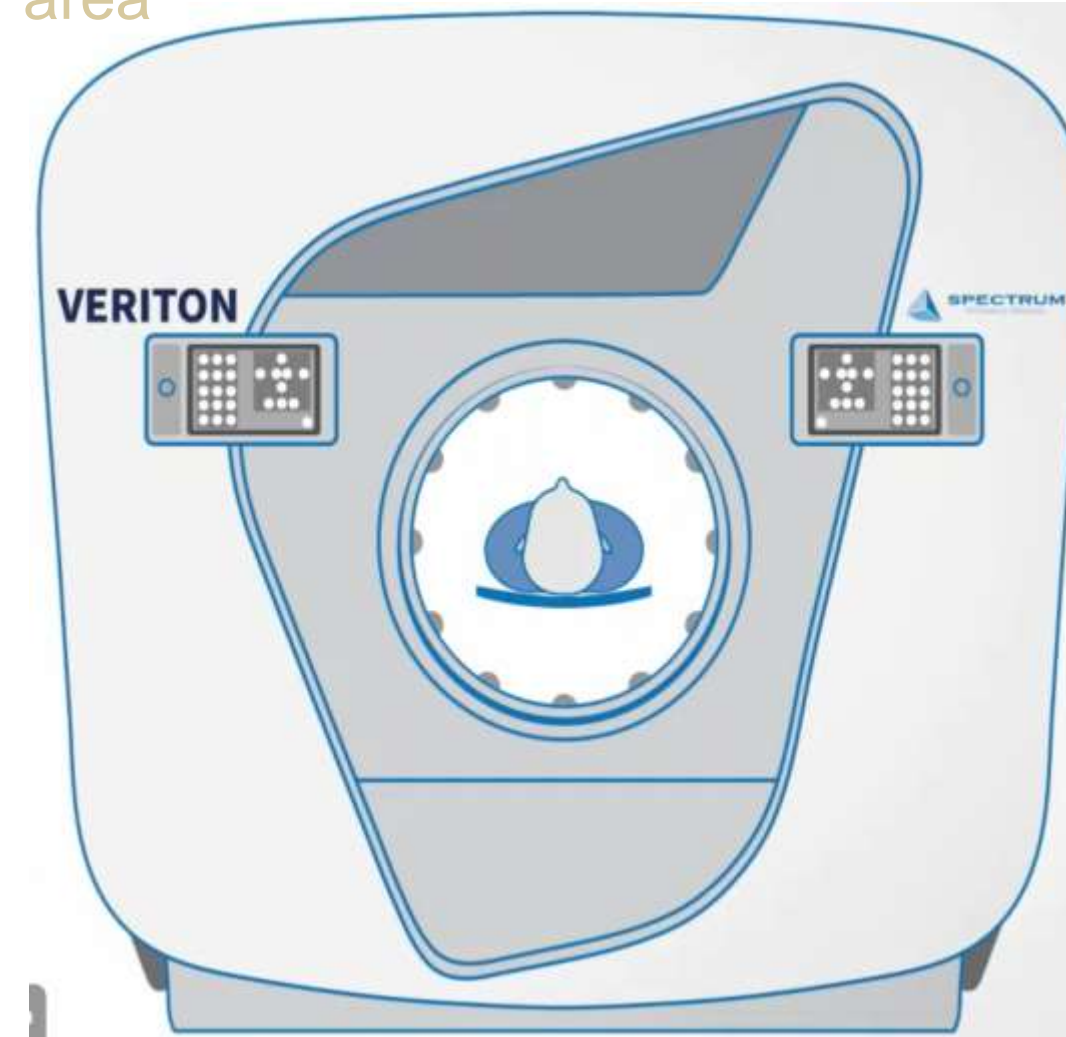


Conventional Dual-Head

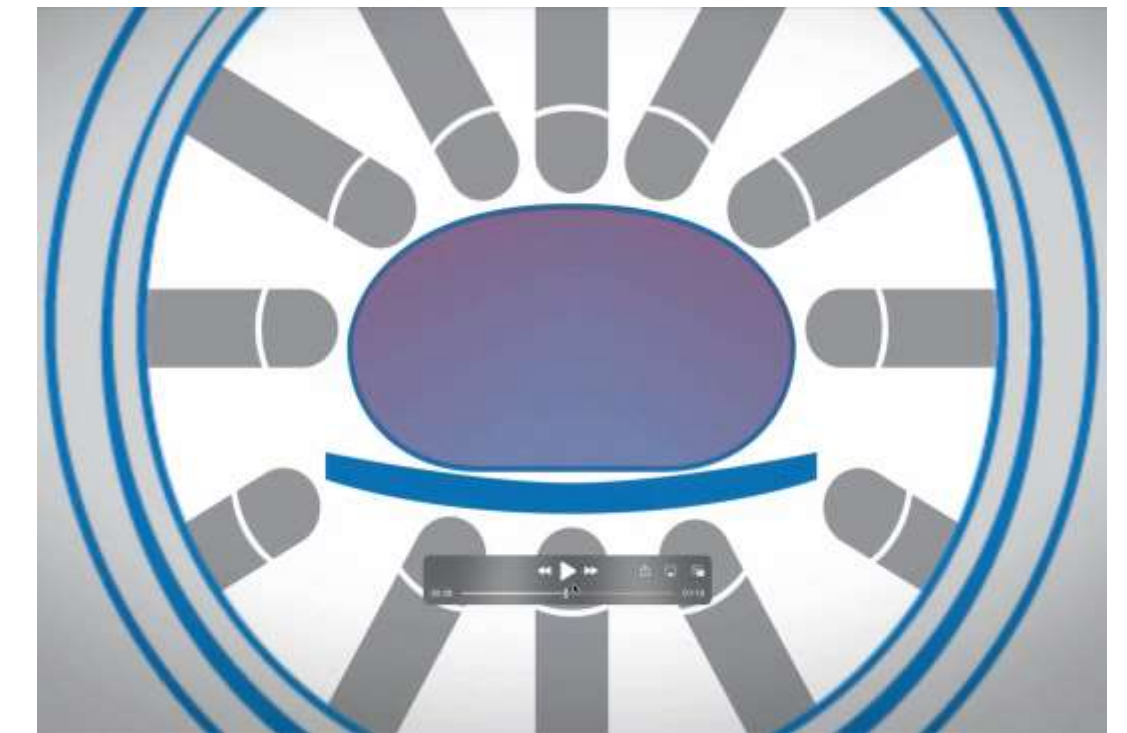
Unused detector area



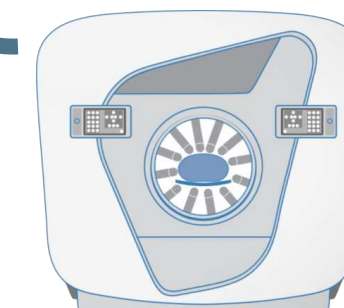
Full-ring CZT



Detector Swiveling

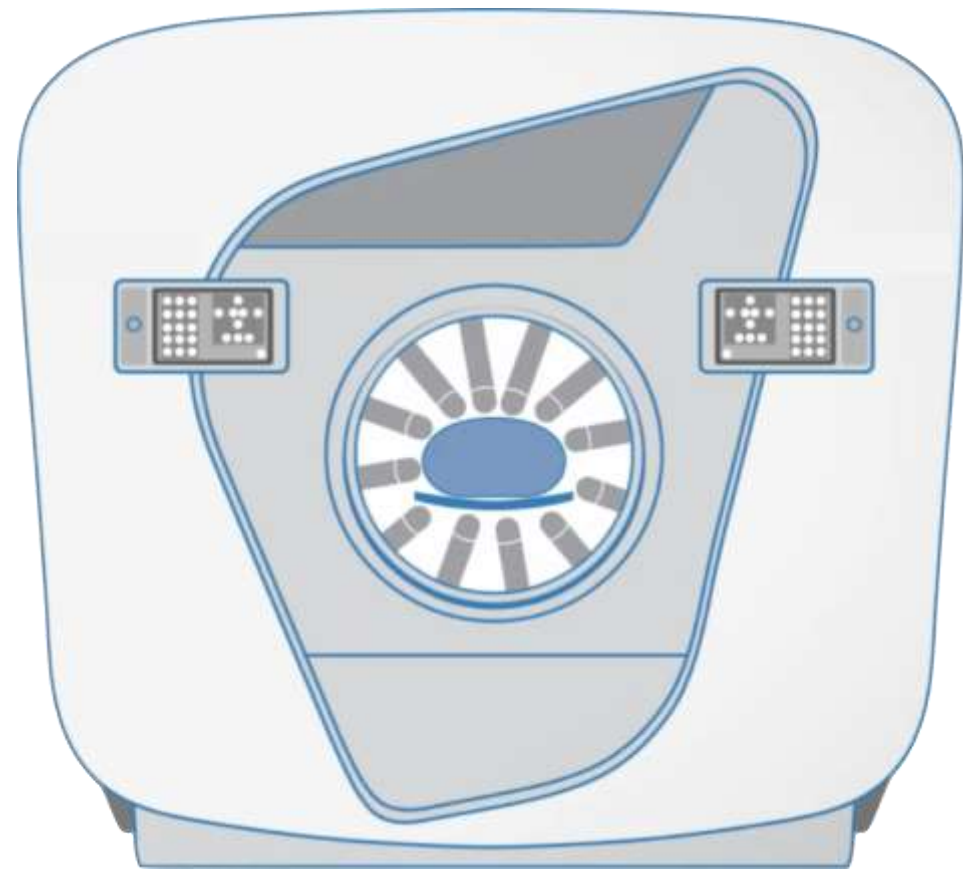


Better Sensitivity and Spatial Resolution because of 360-deg Geometry, Closer-Contouring, and Detector Swiveling/Adaptive Imaging



Improved Imaging Performance

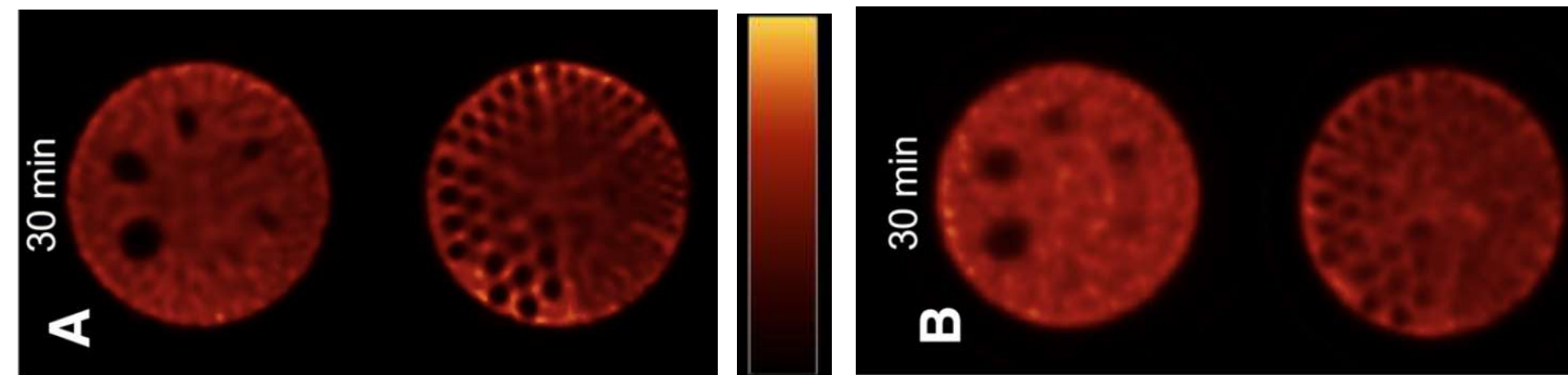
High Sensitivity and Improved Spatial Resolution compared to Conventional Systems



Sensitivity for a ^{99m}Tc point source in head phantom (cps/MBq)

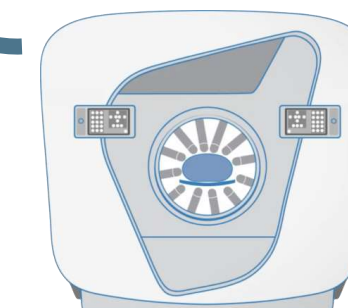
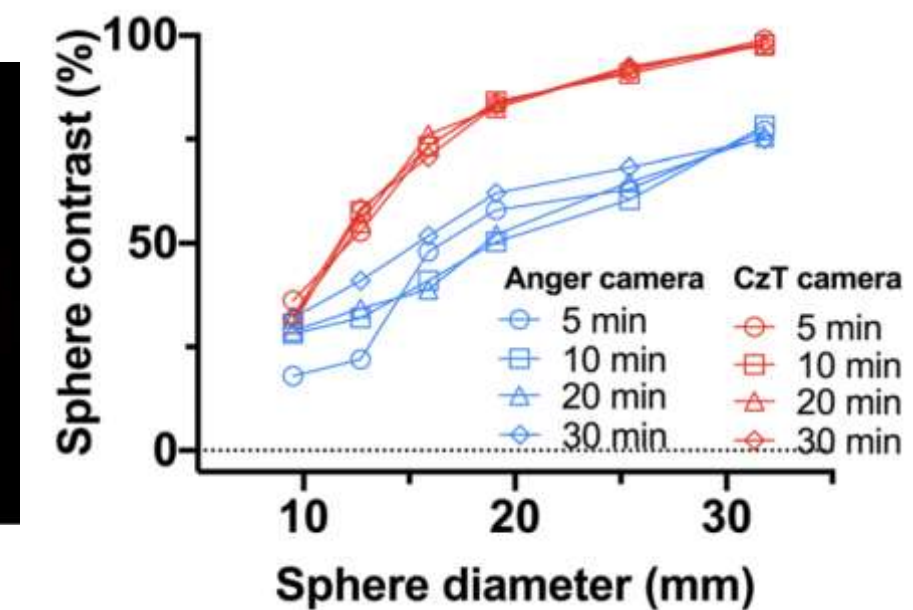
Conventional dual-head LEHR	57.0
Full-Ring CZT (Non-Focus Mode)	73.4 (+22%)
Full-Ring CZT (Focus Mode)	342.4 (+500%)

Tomographic Spatial Resolution



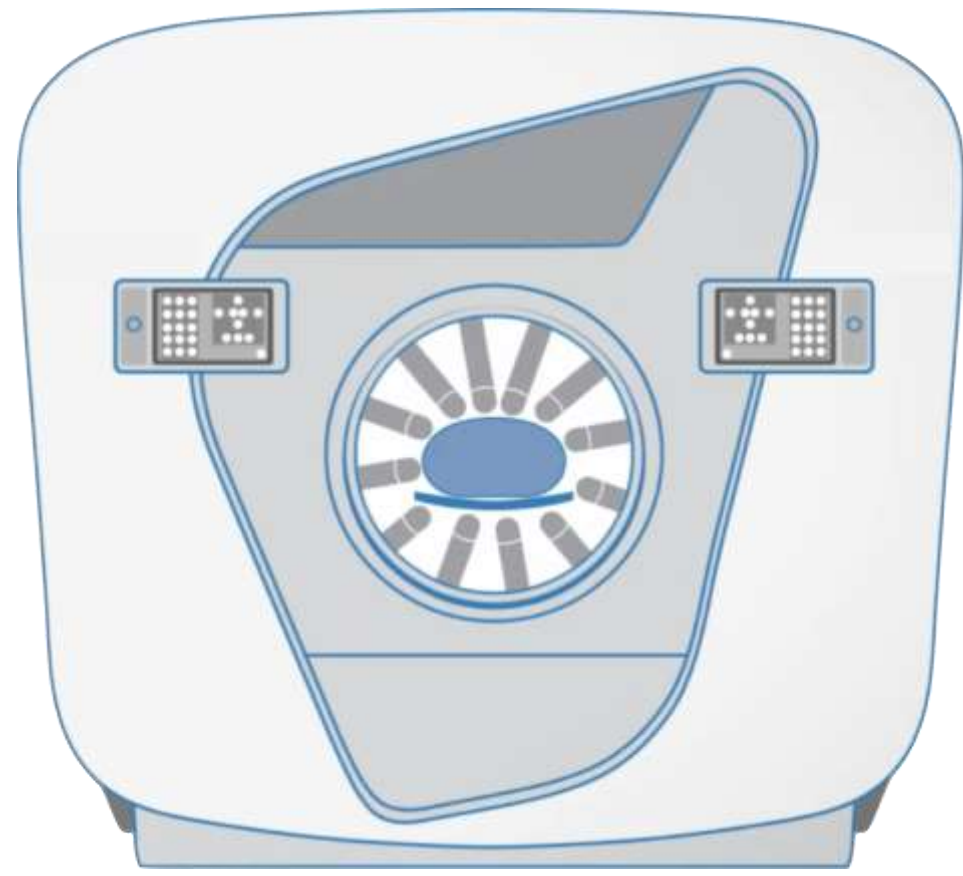
Full-Ring CZT

Conventional





FULL-RING CZT SPECT/CT



Improved Imaging Performance

High Sensitivity and Improved Spatial Resolution compared to Conventional Systems

Adaptive Imaging

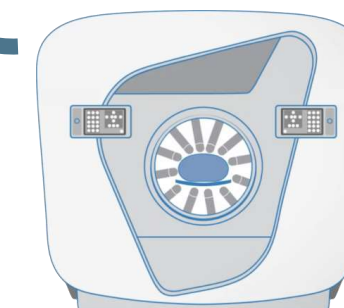
Novel Detector Arrangement
Versatility in Application
Optimum Performance

Quantitation

CT-based Attenuation Correction
Scatter Correction
Sophisticated Reconstruction Algorithm
Built-into the software

3D Dynamic Imaging

360° Acquisition with no Gantry Rotation
High Sensitivity
High Count Rate Capability



3D Dynamic Imaging

- ❑ 360° acquisition with no gantry rotation
- ❑ 3 - 10 sec frame rate
- ❑ High Count-Rate capability

ATTR-Cardiac Amyloidosis 3D Dynamic Imaging

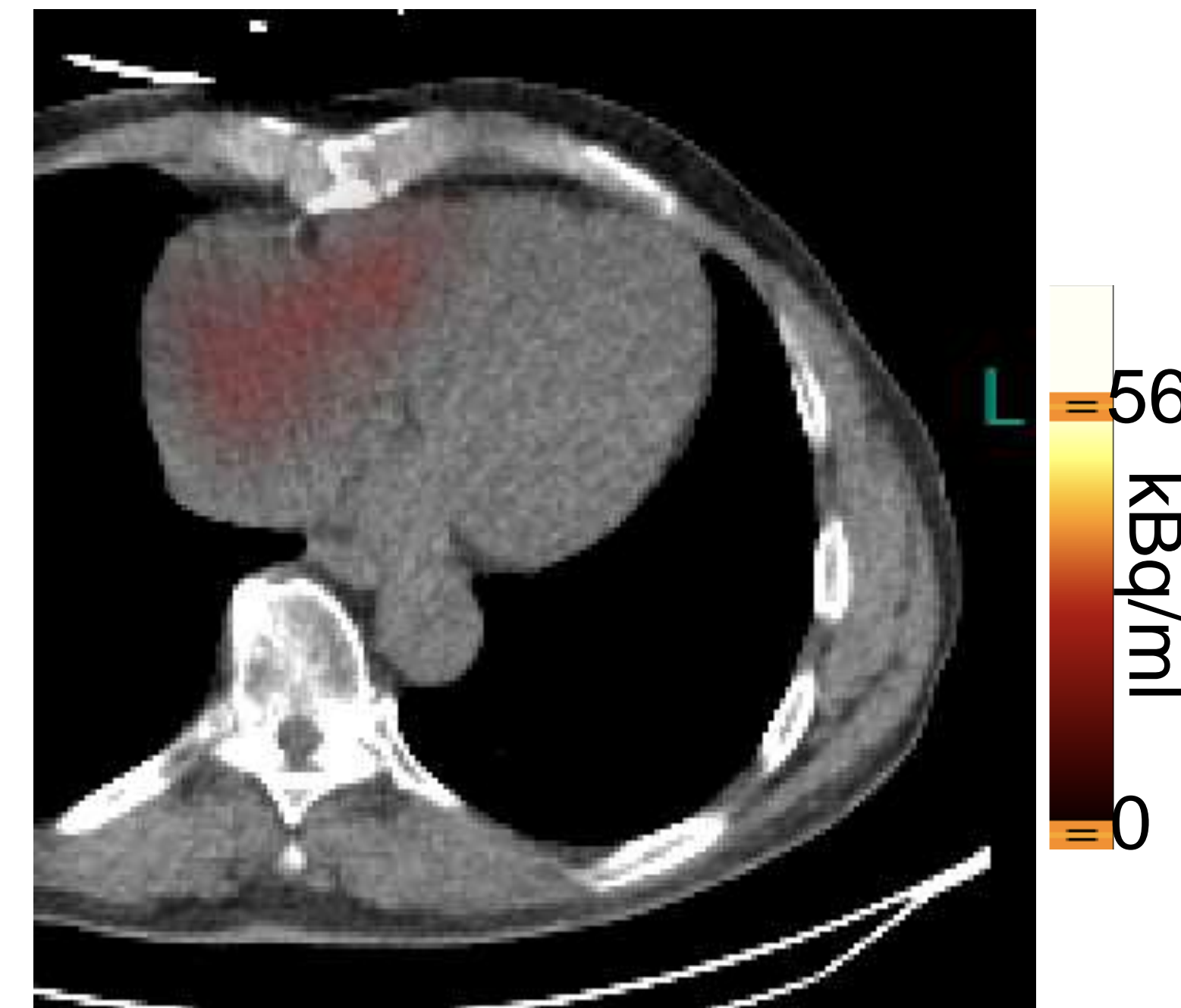
- 20 min dynamic scan
- 138 frames of 8.7 sec
- 20.3 mCi (^{99m}Tc-PYP)

- 71y Male
- BMI of 26.3
- Disease Grade 3

Coronal View

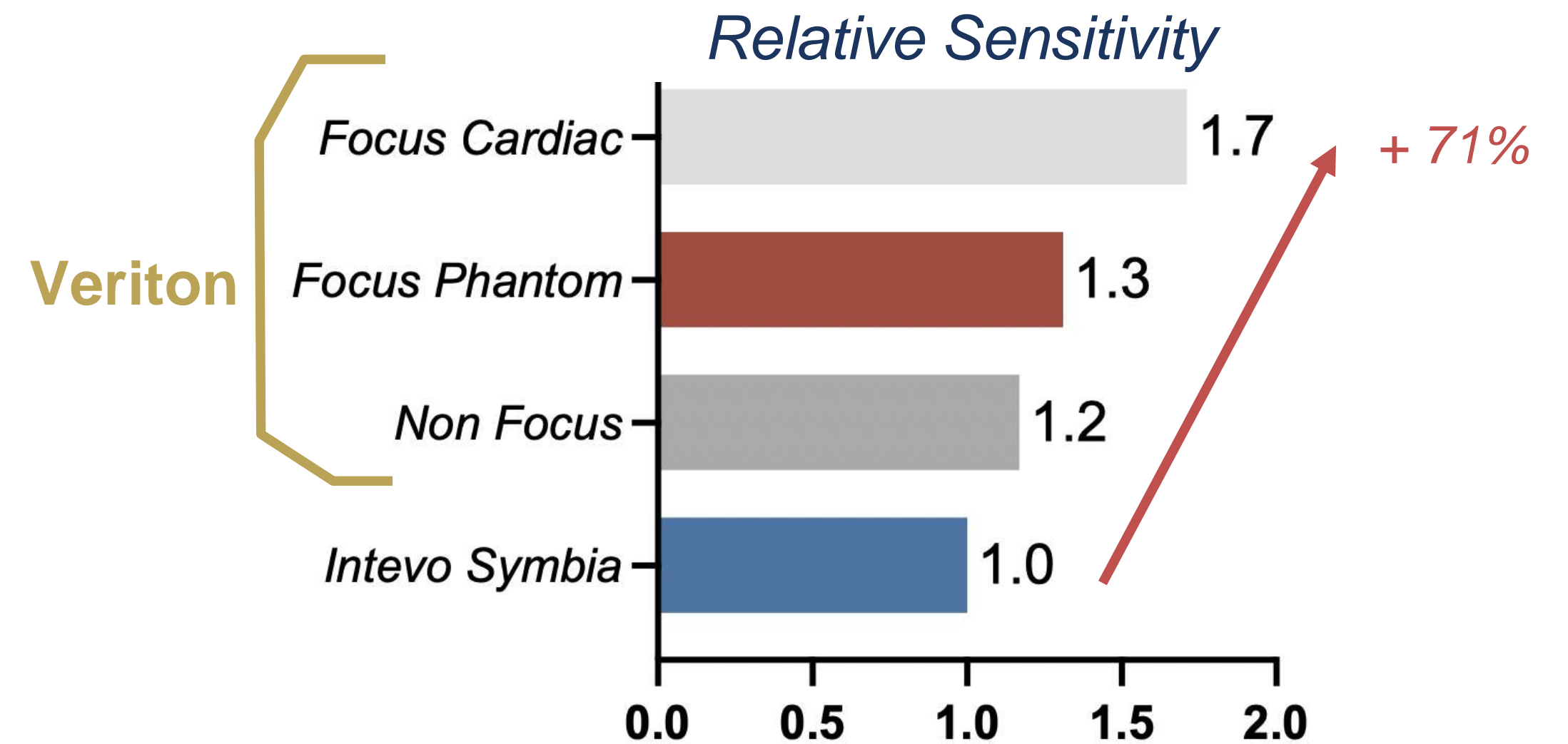
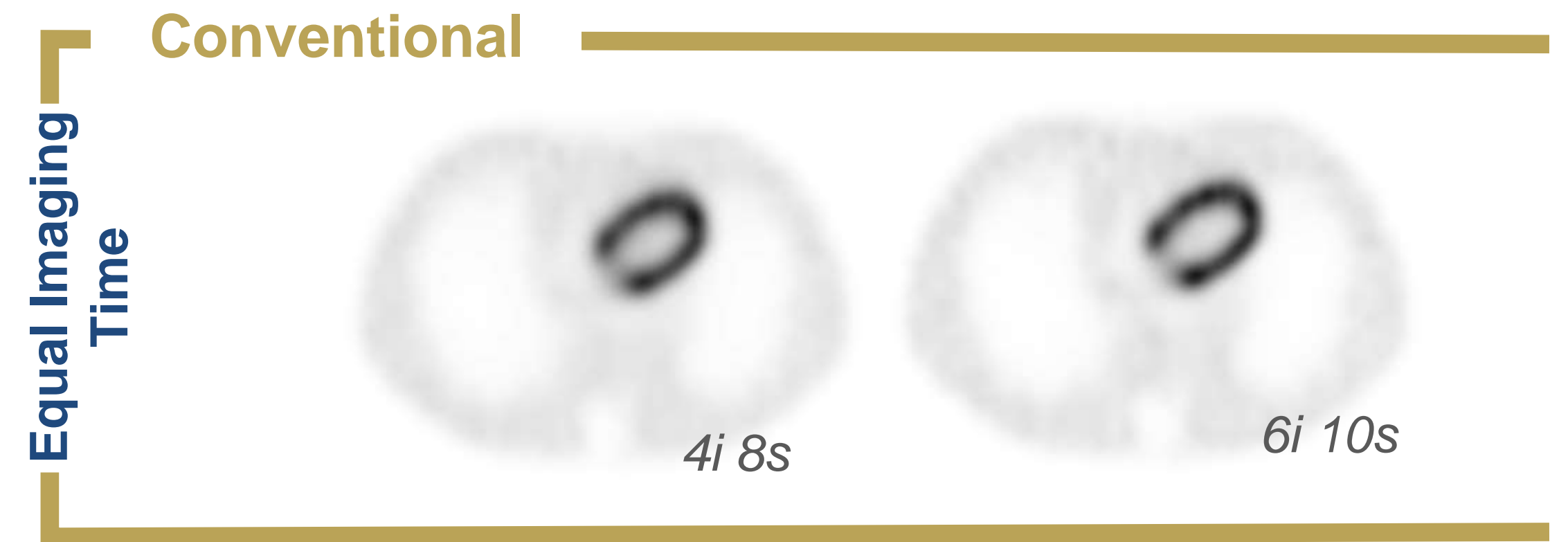
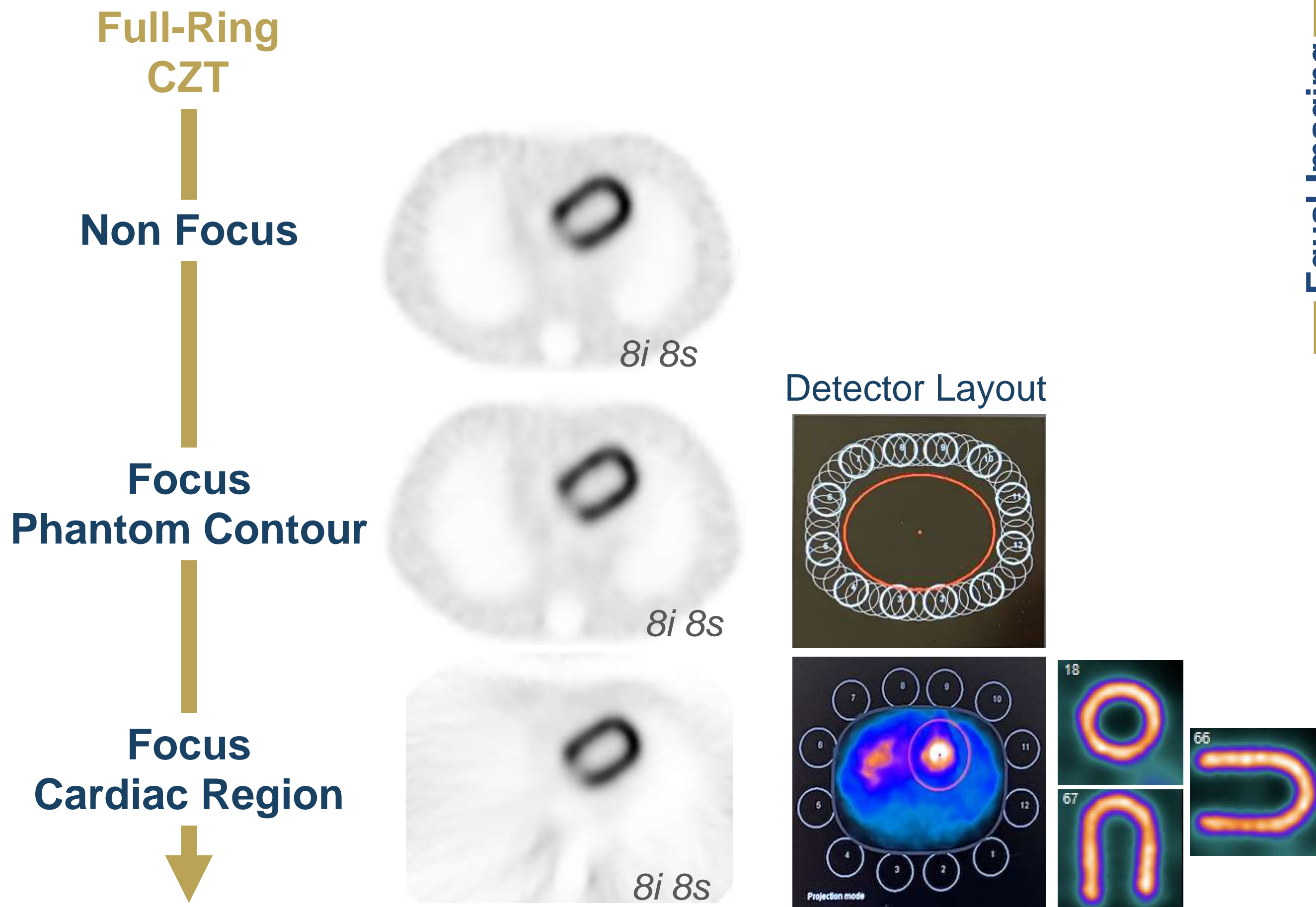


Transaxial View



Full-Ring CZT SPECT/CT has major advantages for MPI,

High sensitivity and spatial resolution can improve qualitatively and quantitatively MPI images compared to conventional SPECT/CT

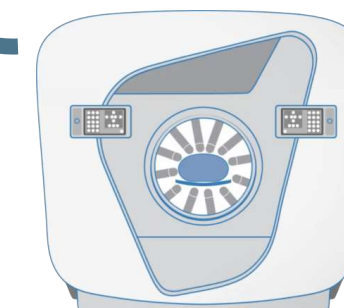




Full-ring CZT SPECT/CT has multiple advantages for MPI,

- High sensitivity and spatial resolution that can enhance qualitatively and quantitatively MPI images compared to conventional SPECT/CT and cardiac CZT SPECT
- Enables accurate Attenuation Correction (AC) that can improve accuracy and imaging time (*stress-only imaging*) compared to cardiac CZT SPECT [1-3]
- Can offer Myocardial Blood Flow (MBF) and Myocardial Flow Reserve (MFR) quantitation, thanks to 3D dynamic imaging capability
- Provides a diagnostic CT for coronary calcium scoring and CT angiography studies [4]

[1] Gowd, et al. J. Nucl. Cardiol. 2014
[2] Huang, et al. J. Nucl. Med. 2016
[3] Hendel, et al. J. Nucl. Med. 2002
[4] Slomka, et al. J. Nucl. Med. 2019



AI-Based Attenuation Correction

Attenuation can cause artefacts

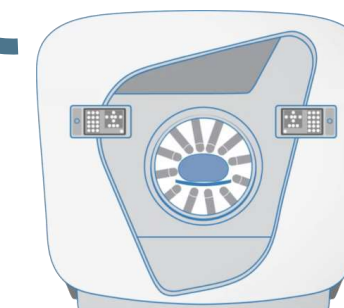
Dual bed positions (**upright/supine**) helps resolving such artefacts, thus reducing the need for rest imaging [1]

AI-based Attenuation Correction has emerged as a powerful tool for low-dose stress-only MPI with single bed position [2-6]

TruCorr[®] is commercially available for D-SPECT [7,8]



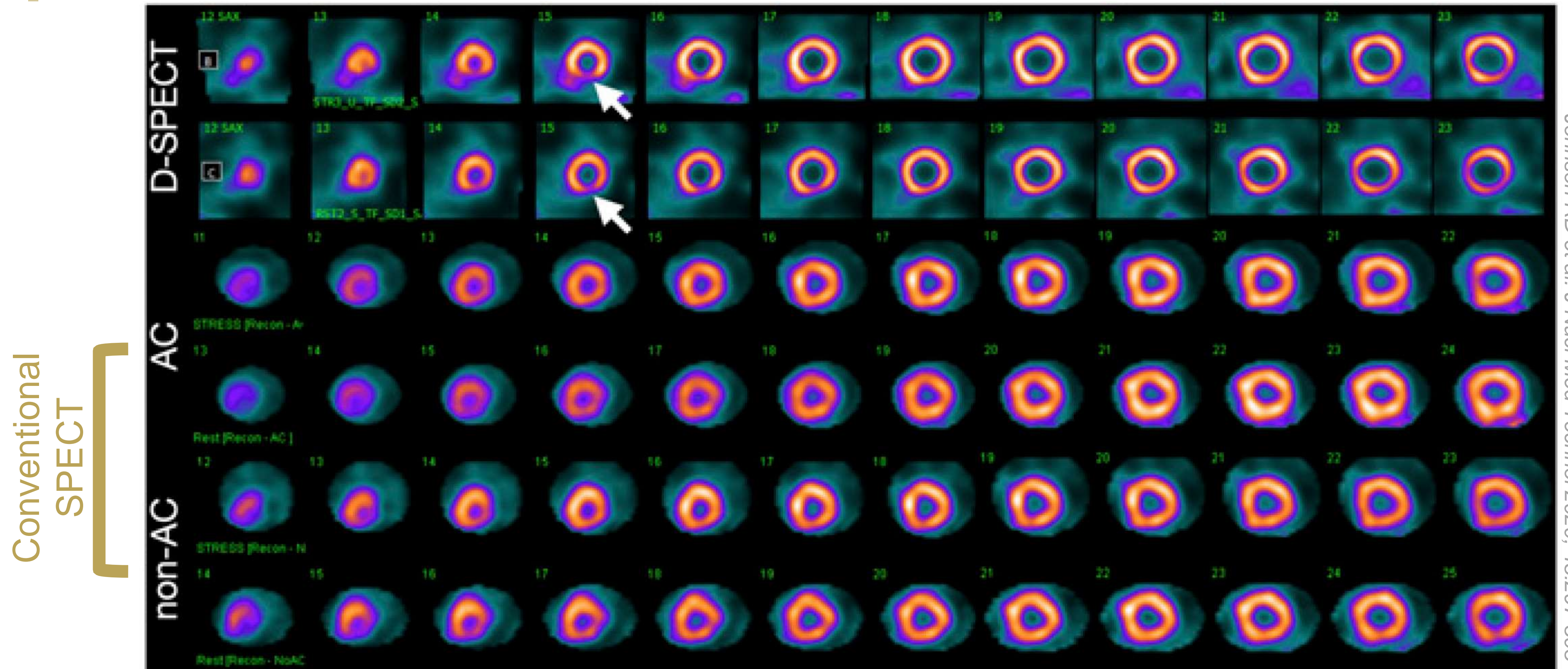
- [1] Nishiyama, et al. Circ. J., 2014
- [2] Yang, et al. J. Nucl. Med., 2021
- [3] Chen, et al. J. Nucl. Cardiol., 2022
- [4] Shi, et al. Eur. J. Nucl. Med. Mol. Imaging, 2020
- [5] Liu, et al. J. Nucl. Cardiol., 2021
- [6] Shanbhaq, et al. J. Nucl. Med., 2022
- [7] Sanchez, et al. Eur. J. Nucl. Med. Mol. Imaging, 2022
- [8] Ochoa, et al. Eur. J. Nucl. Med. Mol. Imaging, 2022



Cardiac CZT SPECT (*DSPECT*) provides,

High Performance Cardiac Imaging with Enhanced Image Quality

Up to 8 times Sensitivity Increases compared to Conventional Systems

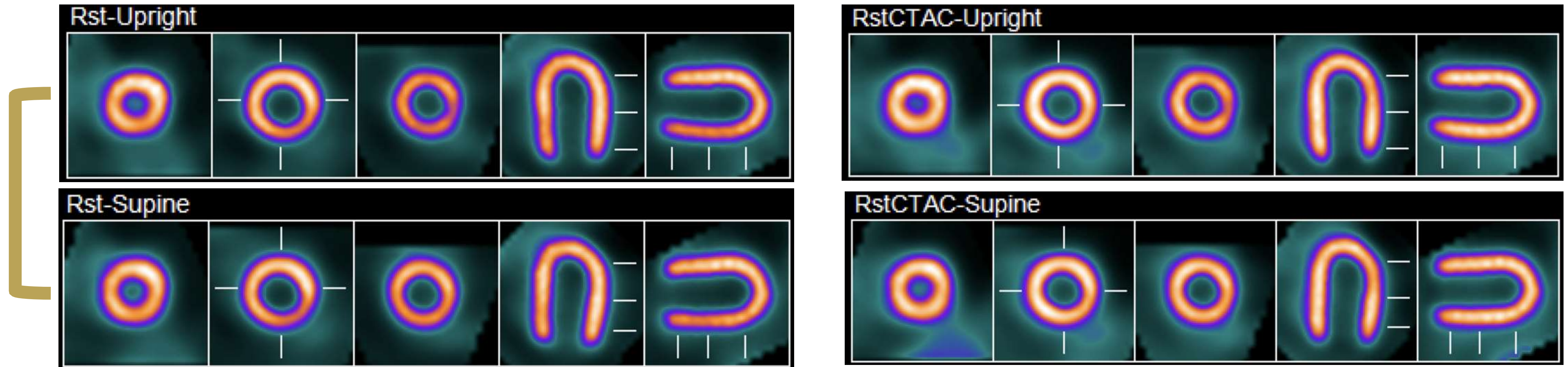


Johnson RD et al. J Nucl Med Technol 2020; 48:297-303

Slightly better performance of TruCorr[®] (*AI-based AC*) in Supine position

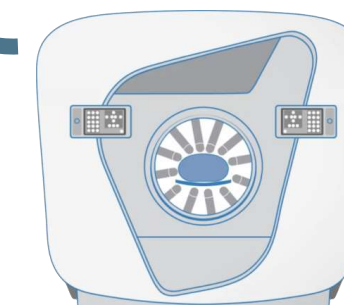
NAC

AI-AC



Torso Phantom Study
 (^{99m}Tc-Sestamibi)

- ❑ Stress equivalent study
- ❑ Activity ratios - Myo:BP:Liver:Bkg 10.9:1.4:3.0:1.0
- ❑ Equal scan time: 5 min 30 sec





MPI WITH CARDIAC CZT SPECT

SPECT Acquisition (^{99m}Tc-Sestamibi)

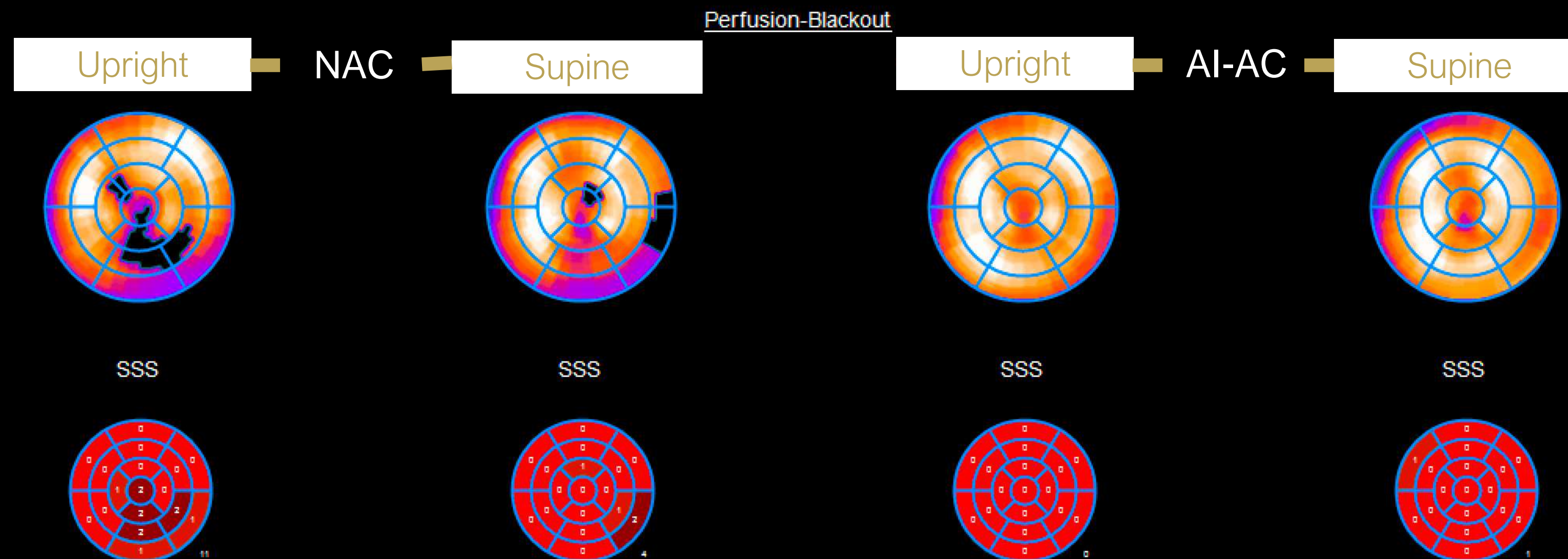
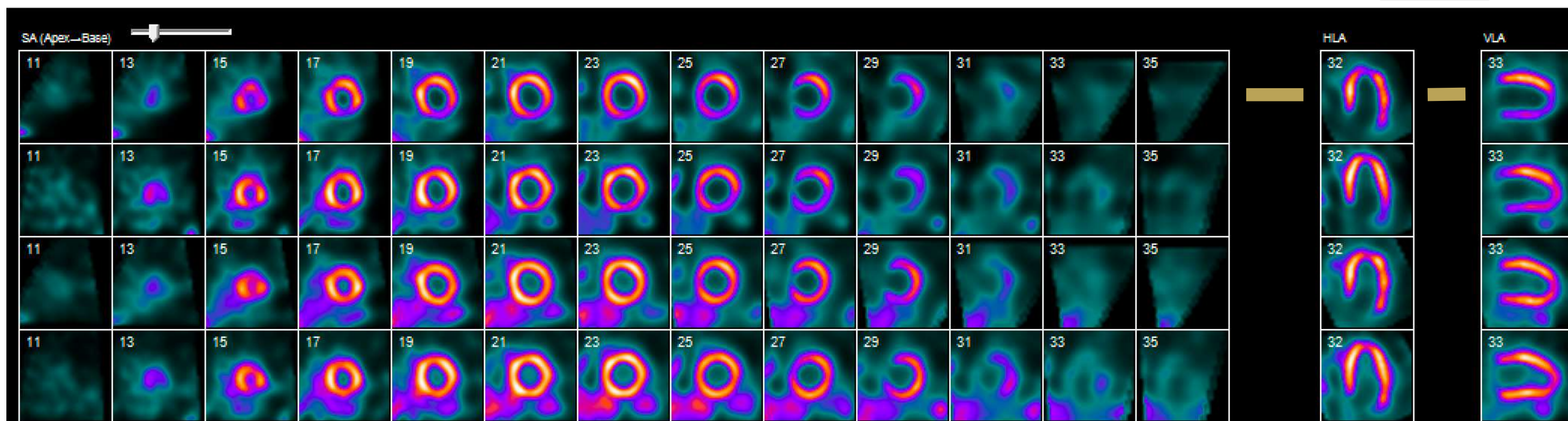
- Stress study
- Dose: 6 mCi
- Scan time: 6 min

- 45y Male
- BMI of 35.6

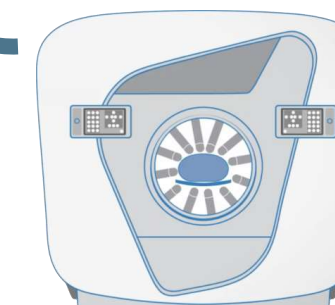
- w/ and w/o AI-based AC
- Supine and Upright

NAC
Upright
Supine

AI-AC
Upright
Supine



0: Normal 1: Mild 2: Moderate 3: Severe 4: Absent



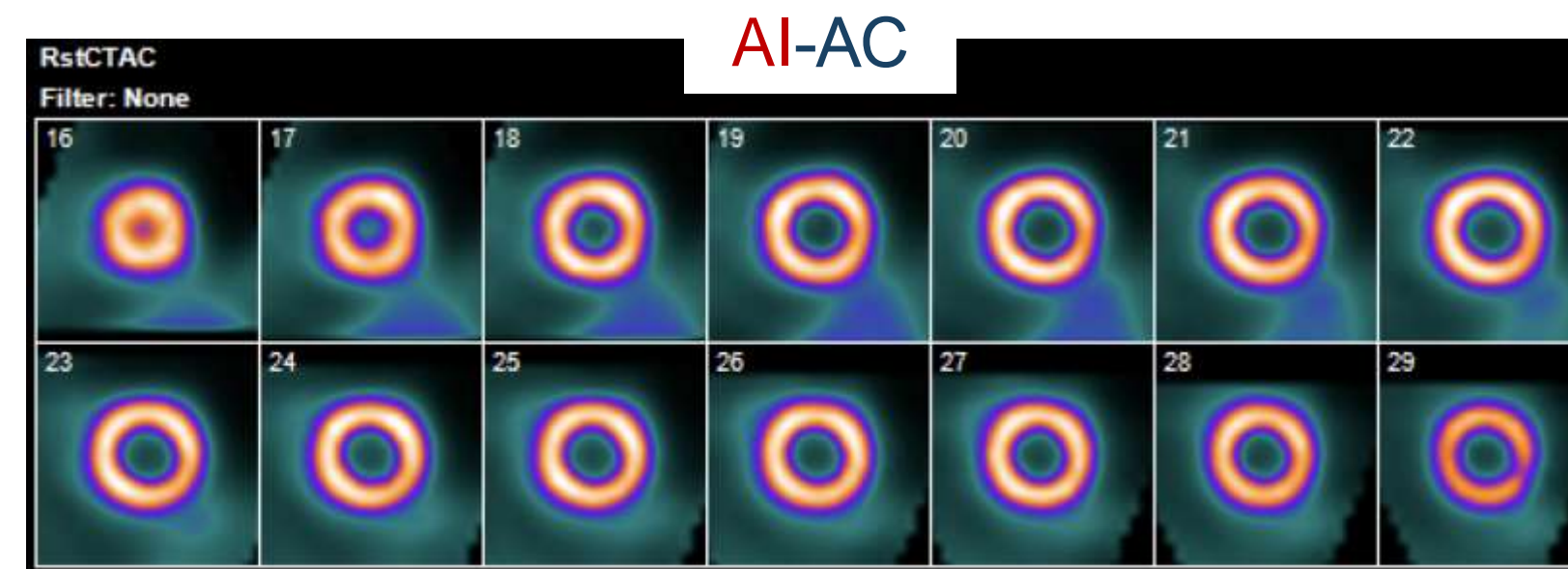
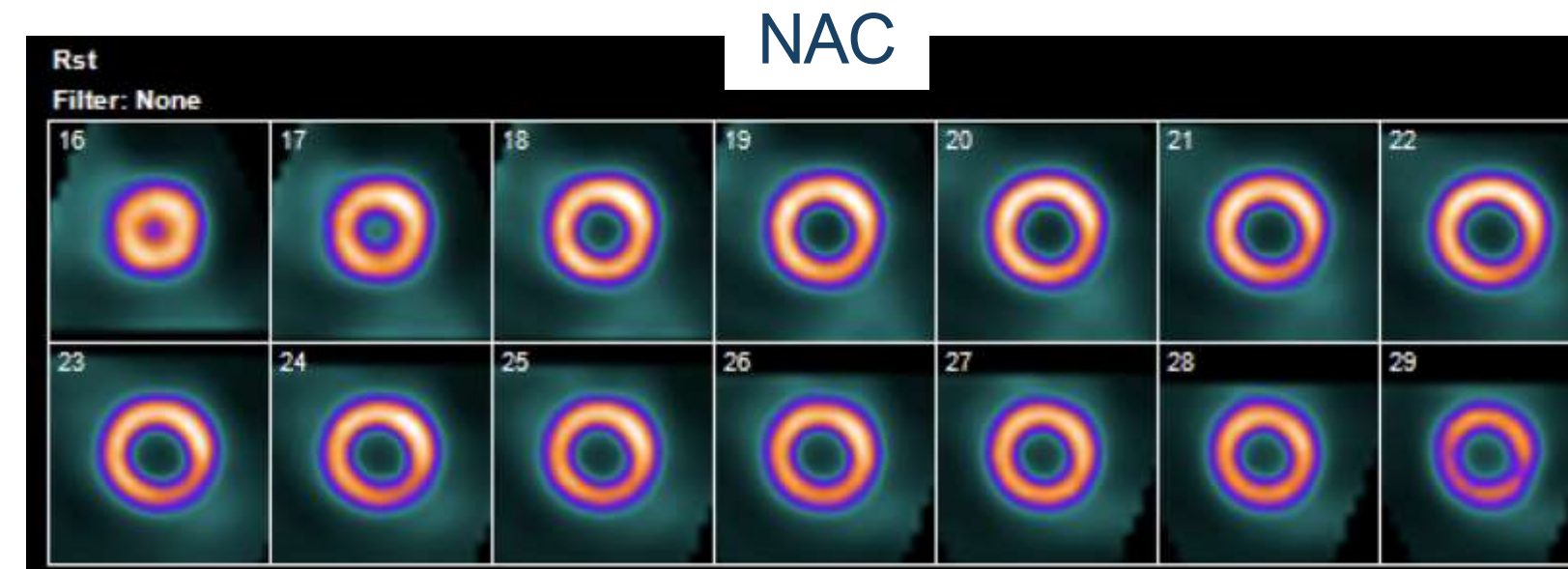


MPI WITH CARDIAC CZT SPECT

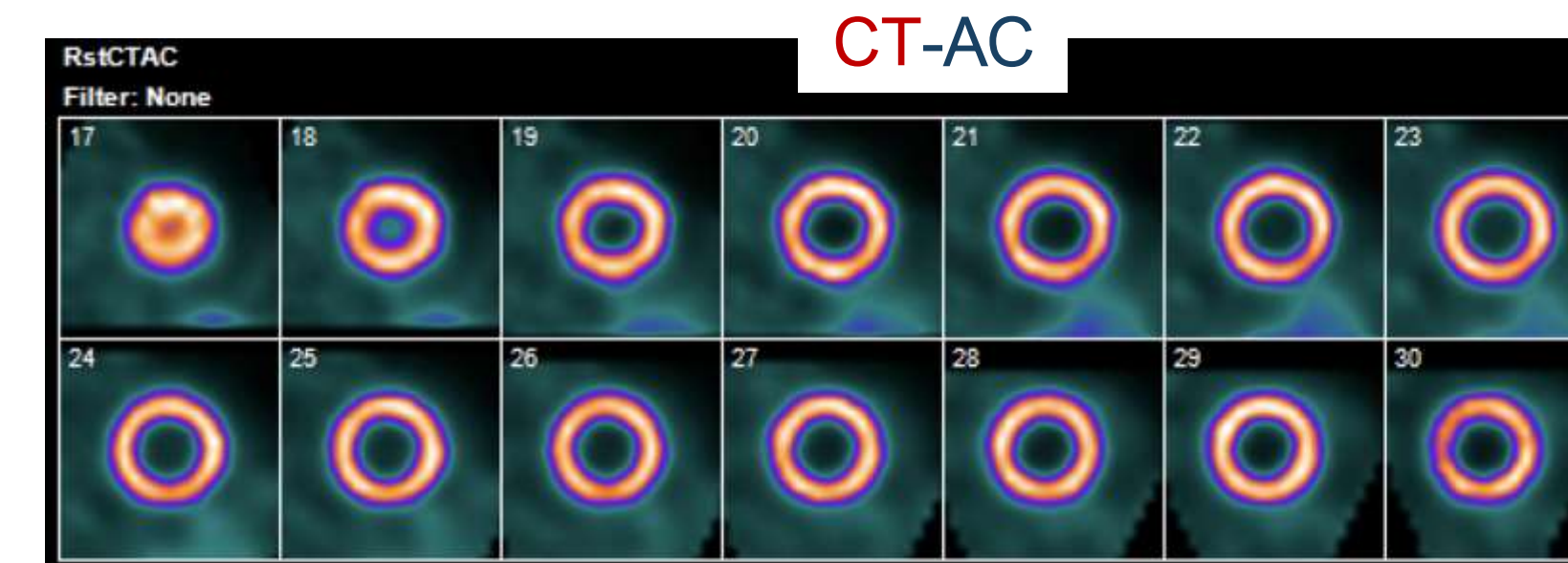
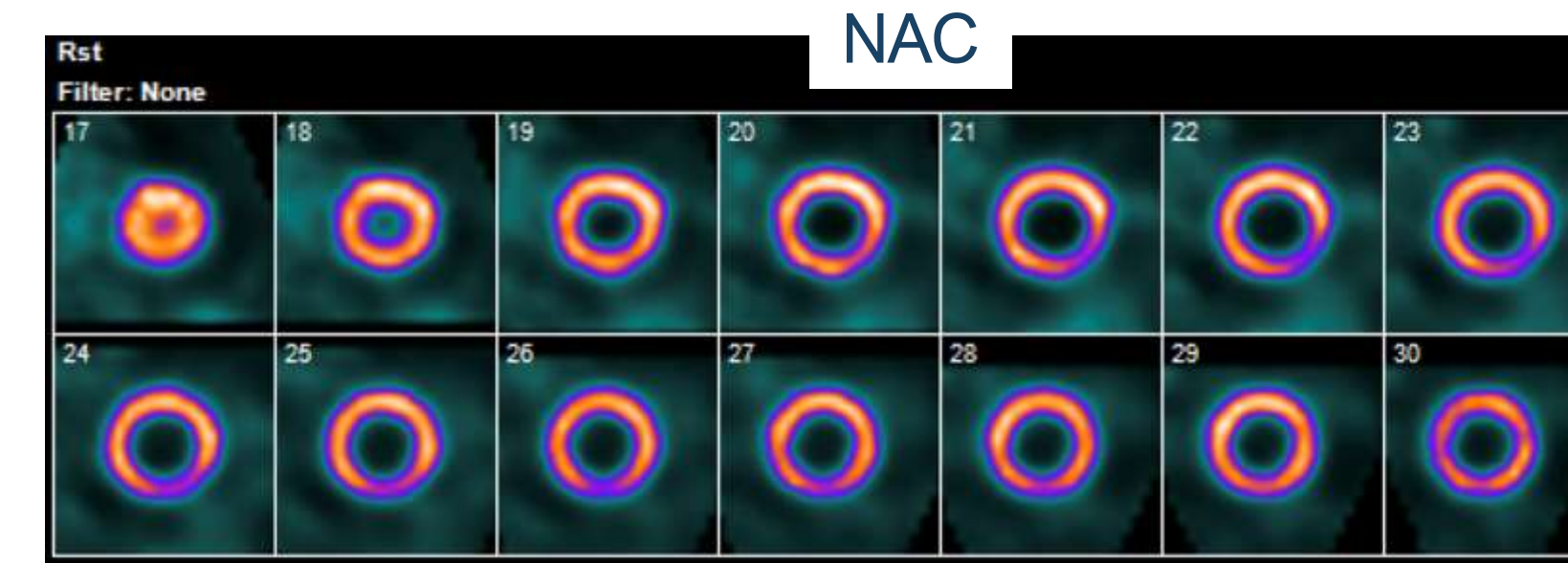
Torso Phantom Study (^{99m}Tc-Sestamibi)

- **Stress** equivalent study
- Activity Ratios –
Myo:BP:Liver:Bkg
10.9:1.4:3.0:1.0
- Equal Scan time: *5 min 30 sec*
- w/ and w/o AI-based AC
- Supine and Upright

Cardiac CZT DSPECT

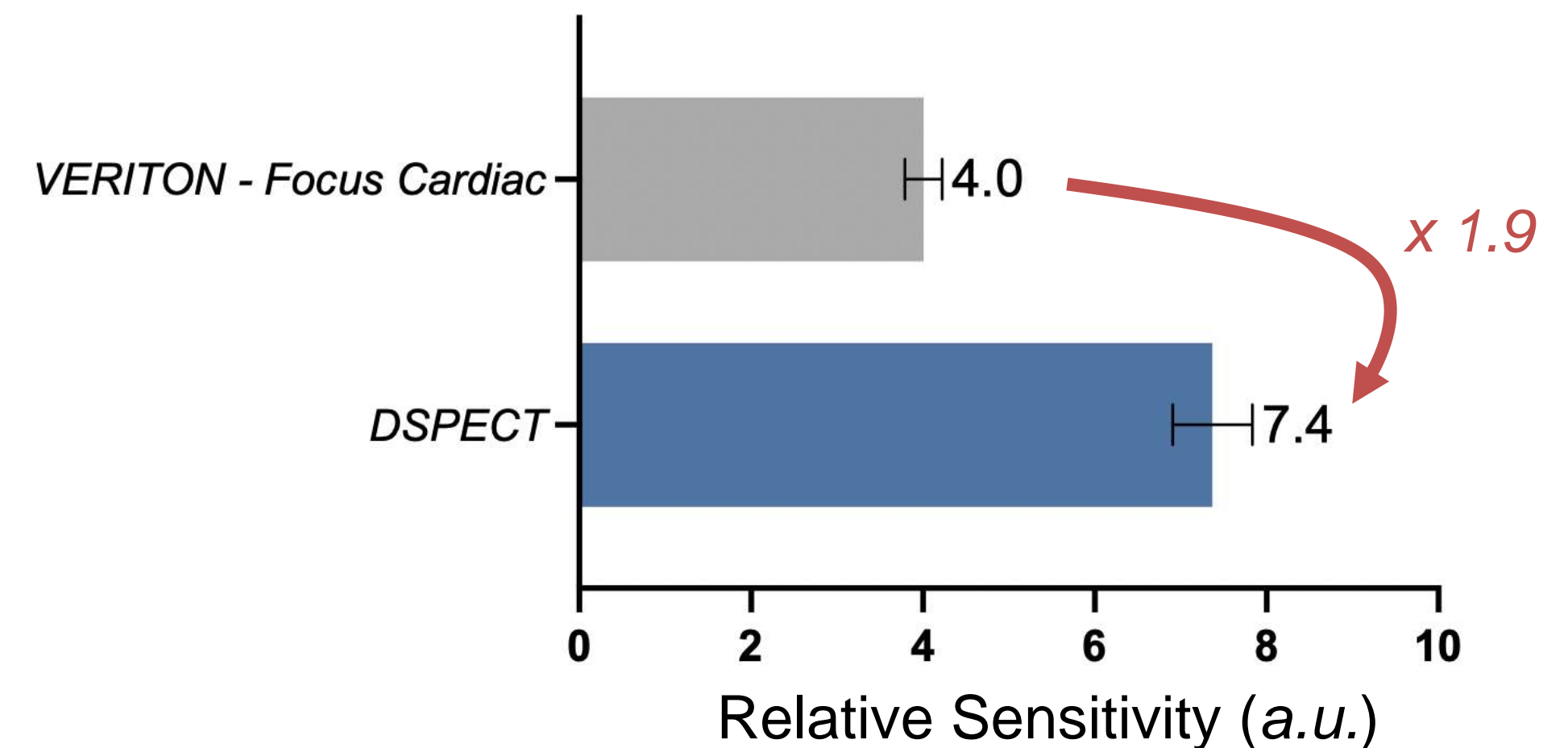


Full-ring CZT Veriton



AI-AC TruCorr[®] works well with phantoms

Full-ring CZT leads to better spatial resolution
at the expense of lower sensitivity





MPI WITH CARDIAC CZT SPECT

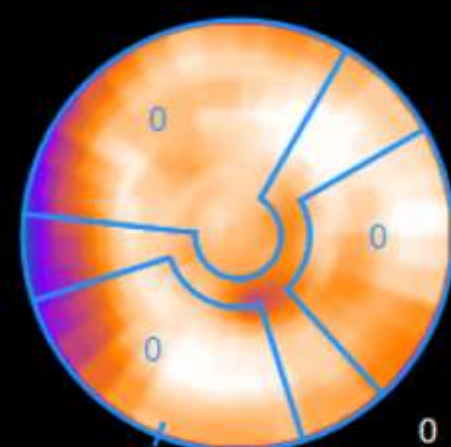
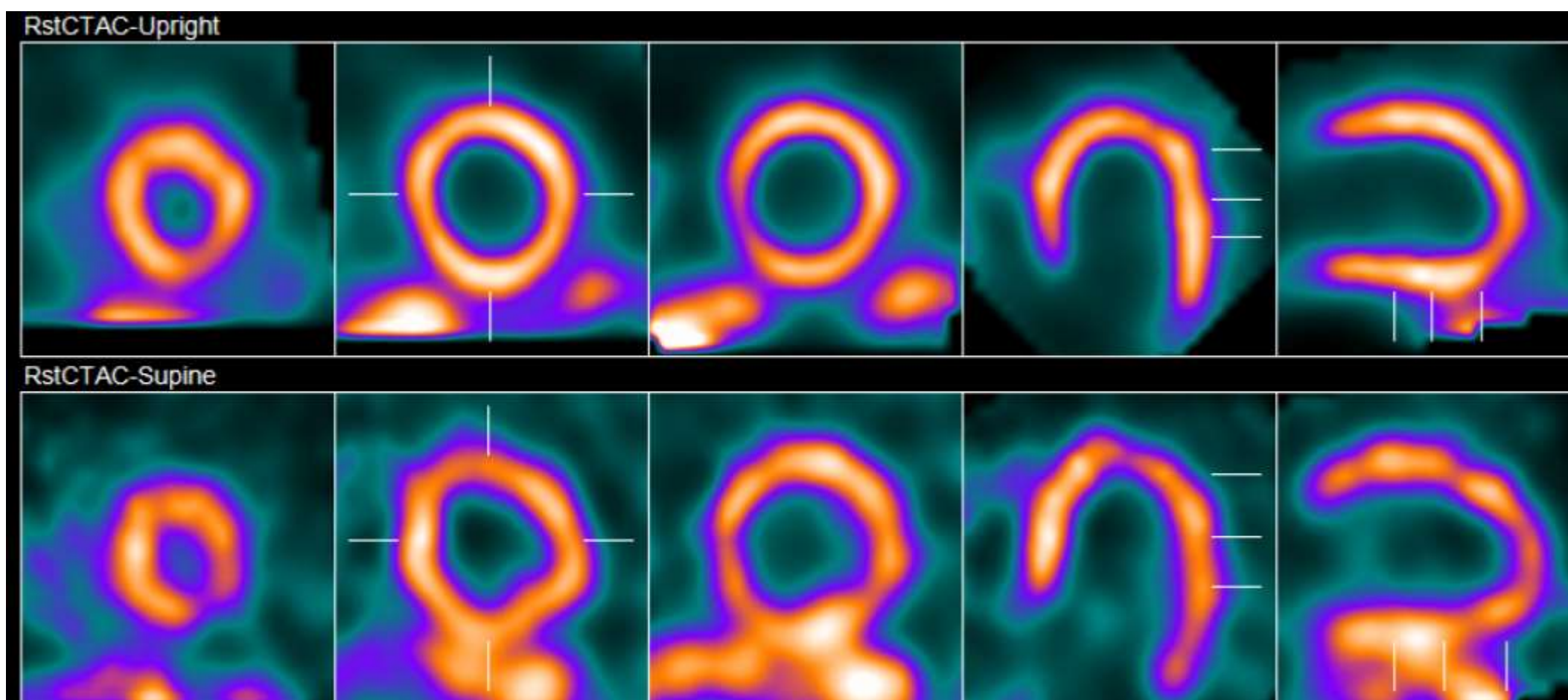
SPECT Acquisition (^{99m}Tc-Sestamibi)

- Rest study
- Dose: 10 mCi
- 68y Female
- BMI of 25.3

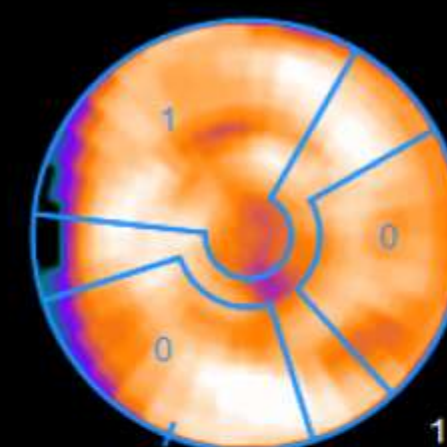
- Veriton CT-AC
Supine
4 min / 3.8 Mcts

- DSPECT AI-AC (90 min delay)
Upright
6 min / 3.4 Mcts

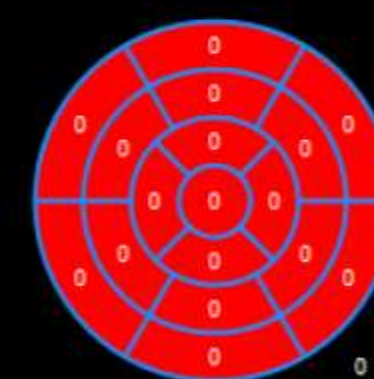
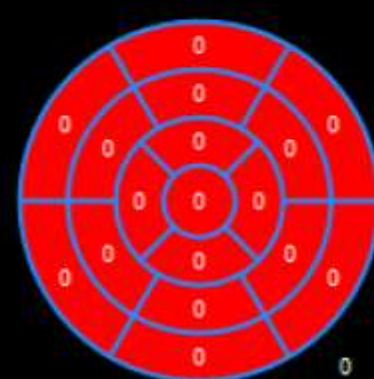
DSPECT AI-AC
Veriton CT-AC



AI-AC

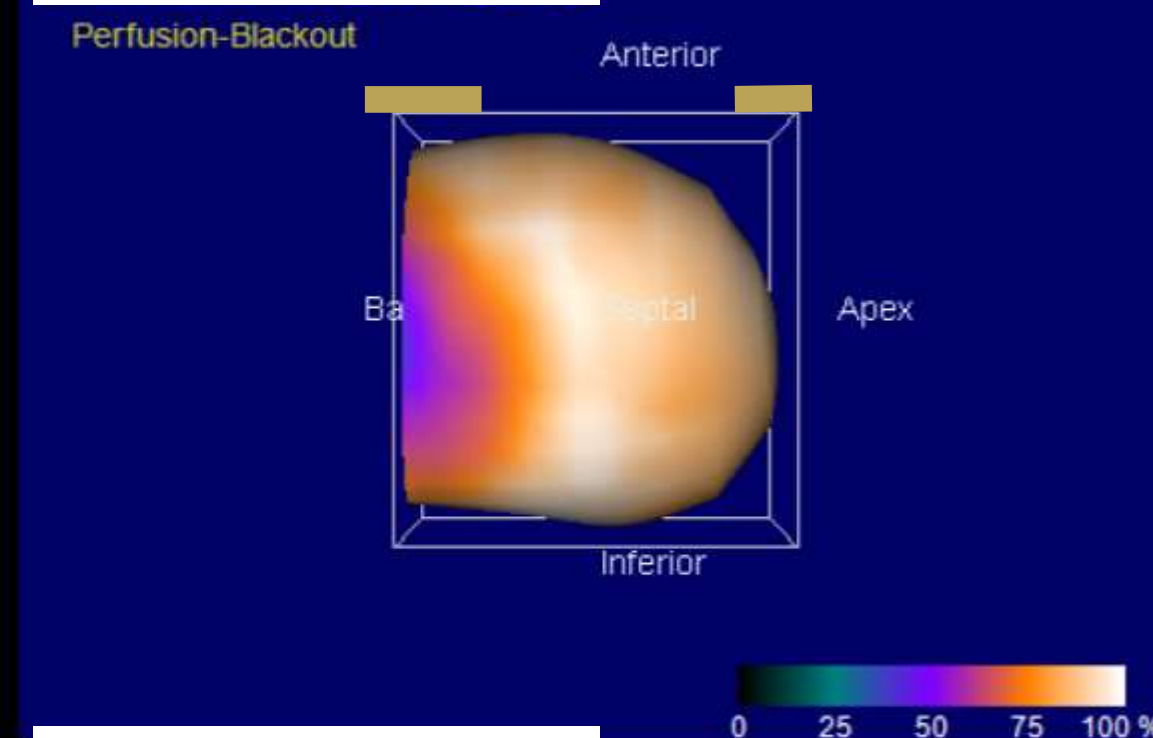


CT-AC

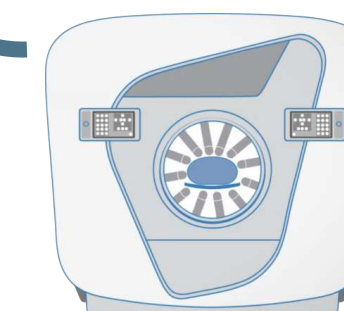
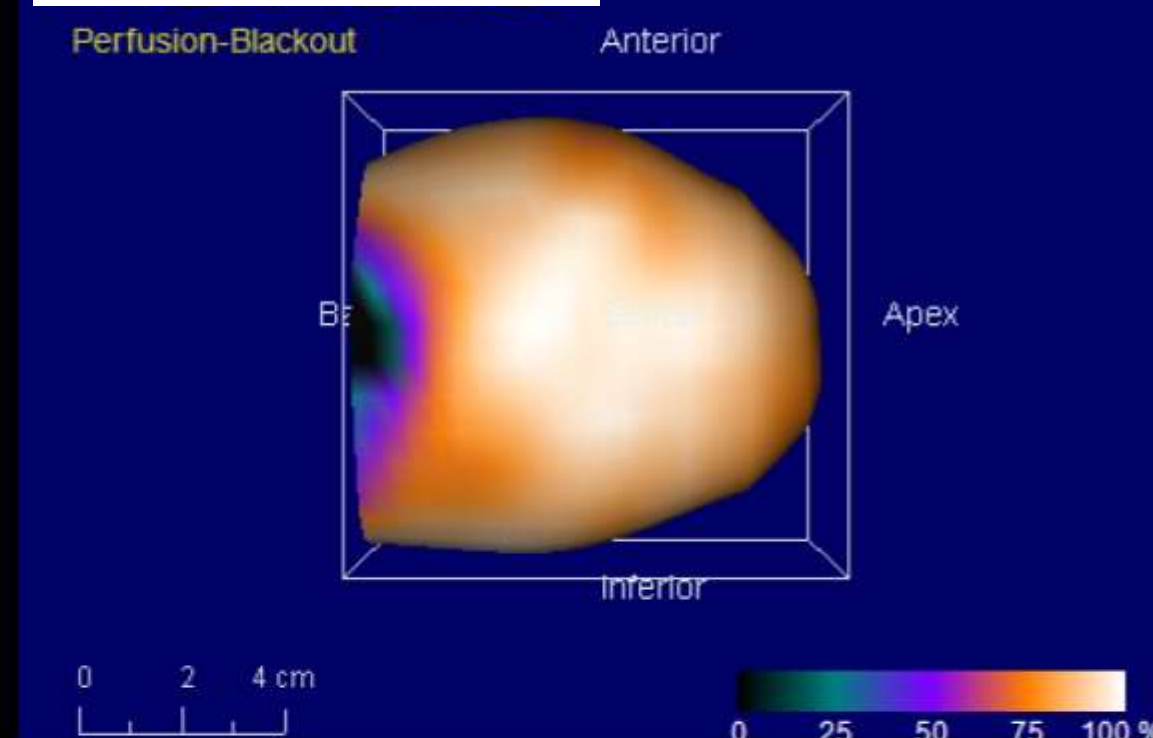


Normal 1: Mild 2: Moderate 3: Severe 4: Absent

D-SPECT AI-AC



Veriton CT-AC



Promising MBF/MFR assessment via SPECT Imaging with Cardiac CZT systems [1-4],

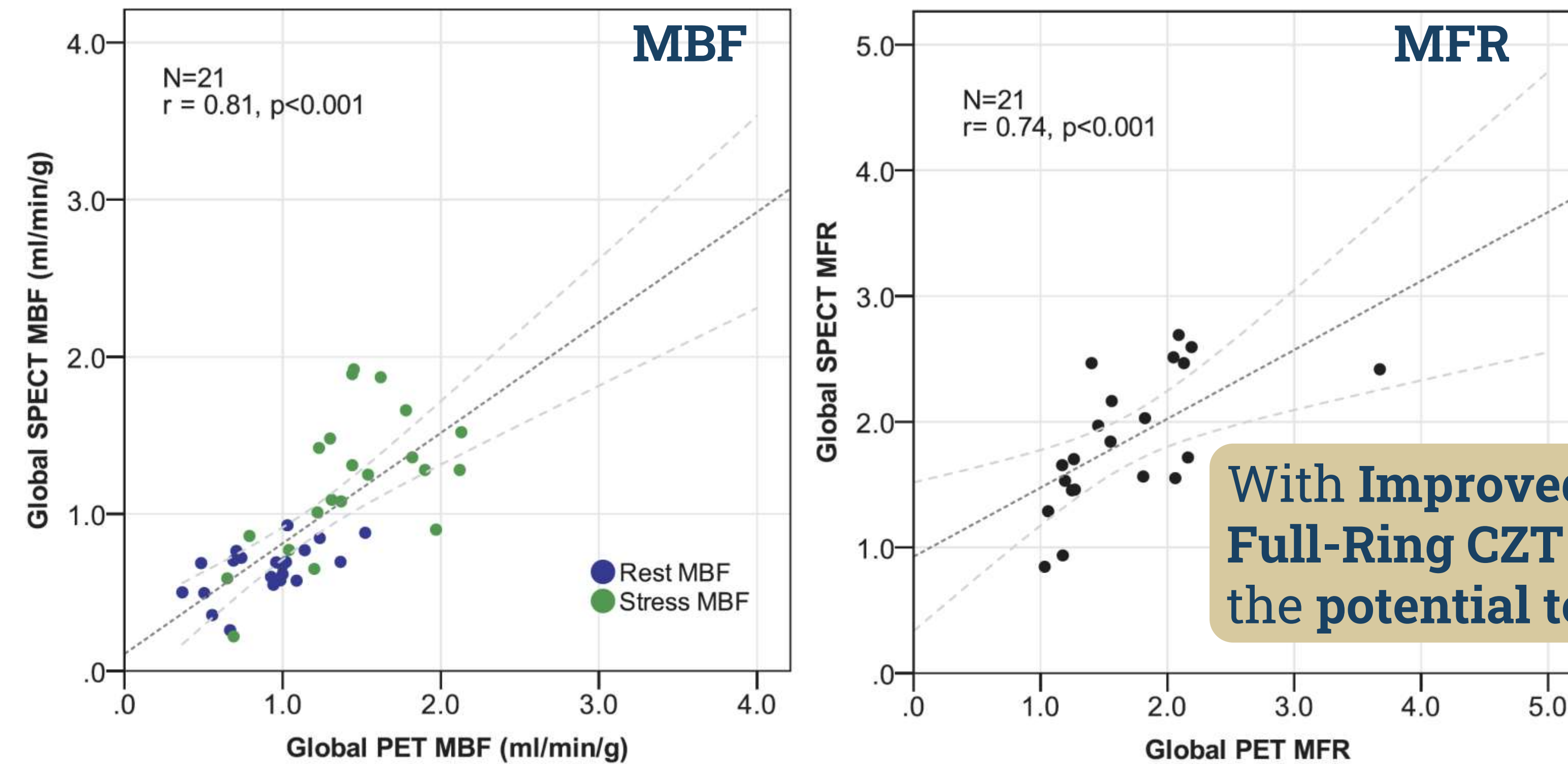
In several single center human studies in which PET was used as reference

[1] Adapted from De Souza, et al. Circulation: Cardiovascular Imaging, 2022

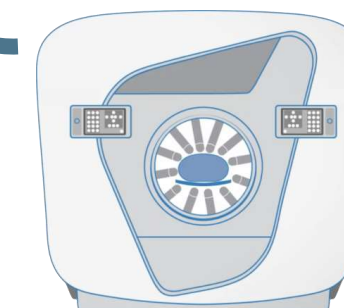
[2] Wells. J. Nucl. Med. 2017

[3] Otaki. J. Nucl. Cardiol. 2021

[4] Agostini. Eur. J. Nucl. Med. Mol. Imaging, 2018



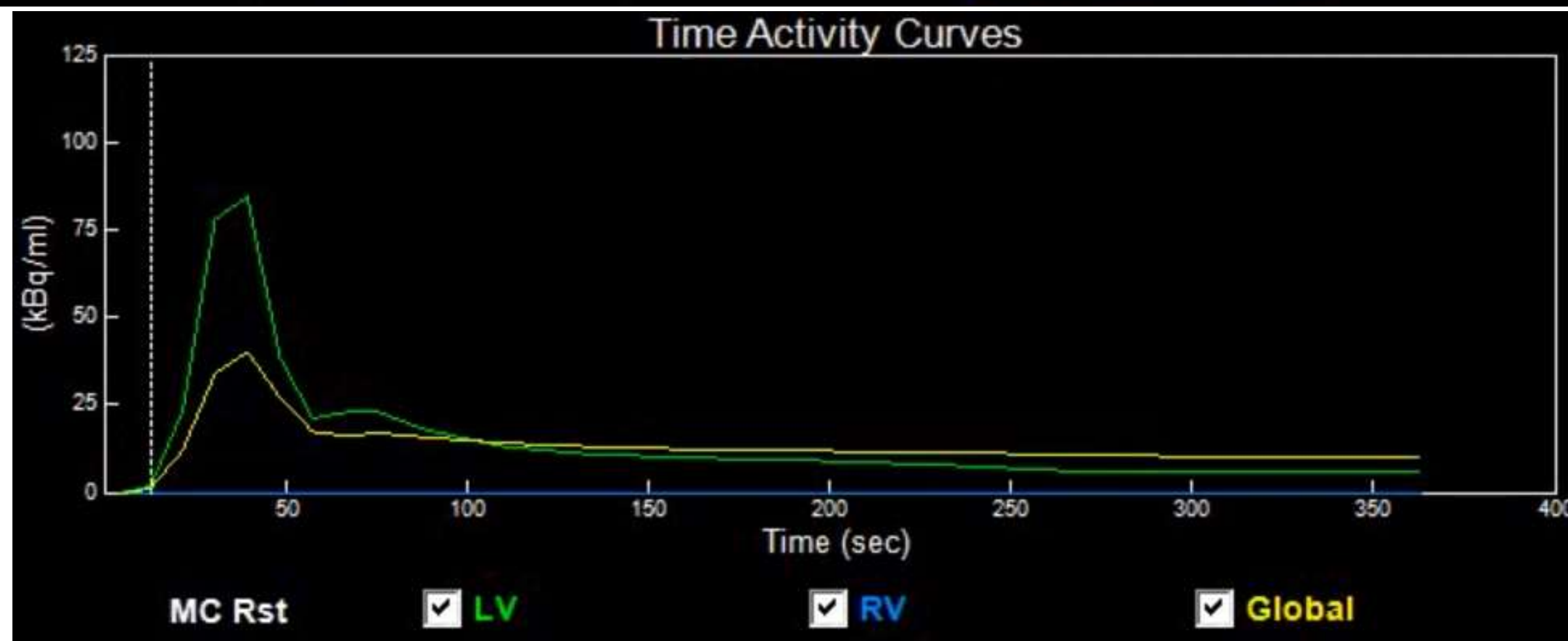
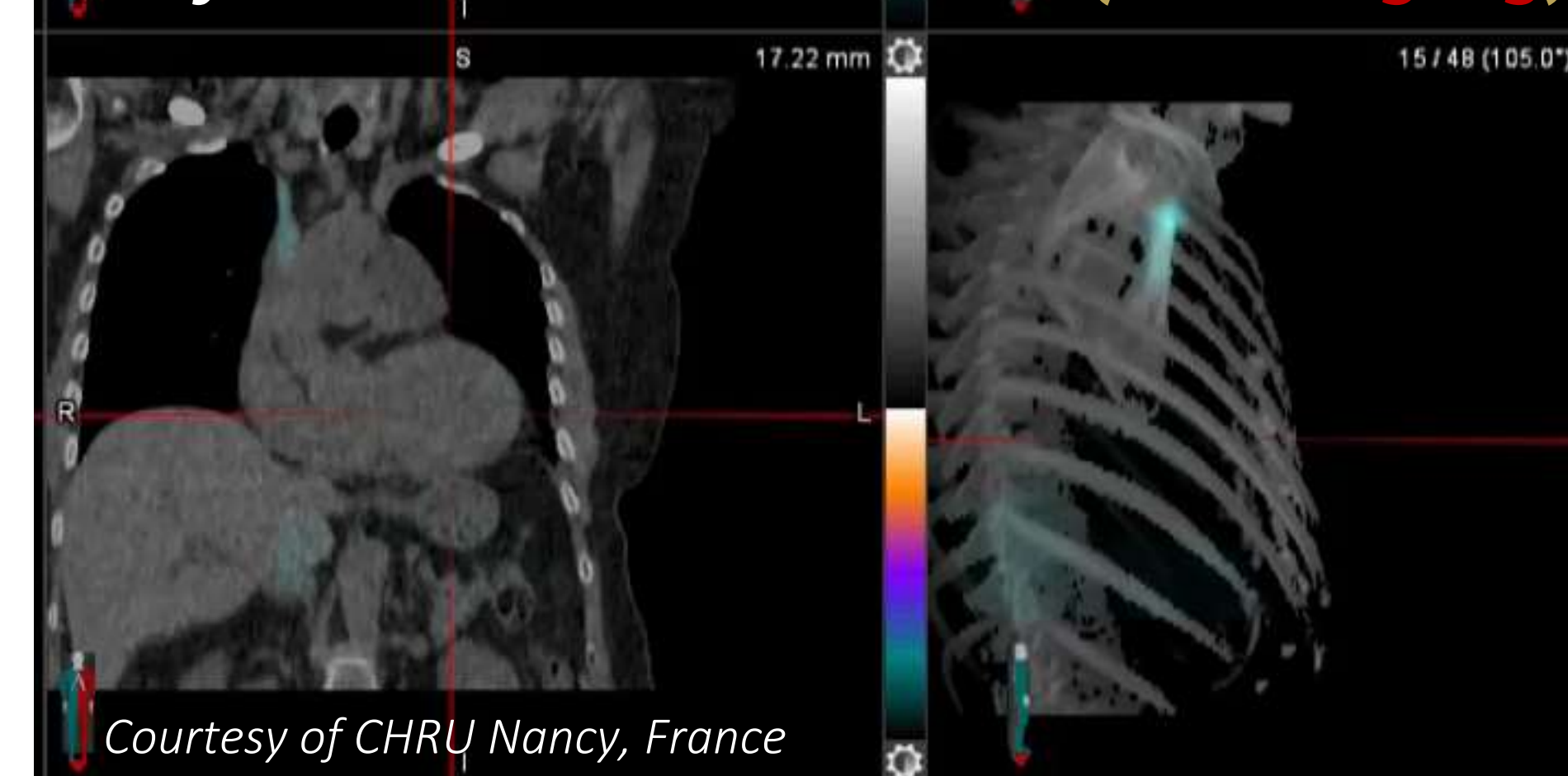
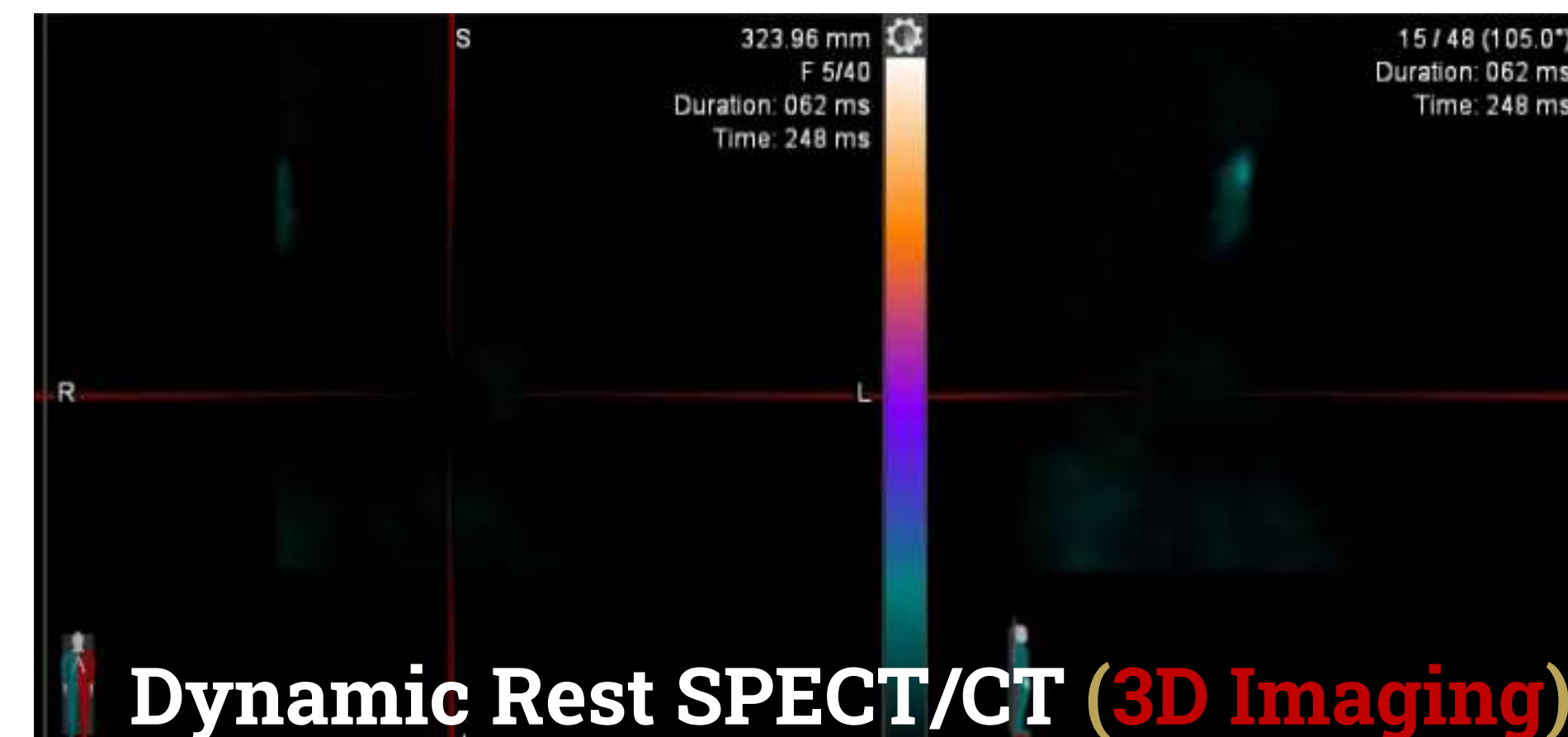
With Improved 3D Dynamic Imaging Capability, Full-Ring CZT SPECT/CT has the potential to enhance further MBF/MFR quantitation.



Full-Ring CZT SPECT/CT has major advantages for MPI,

Offers **Myocardial Blood Flow (MBF)** and **Myocardial Flow Reserve (MFR)** quantitation, thanks to **3D dynamic imaging** capability. CT-AC provides higher accuracy compared to cardiac CZT SPECT

6-min 3D Dynamic SPECT/CT Acquisition (^{99m}Tc-Sestamibi)



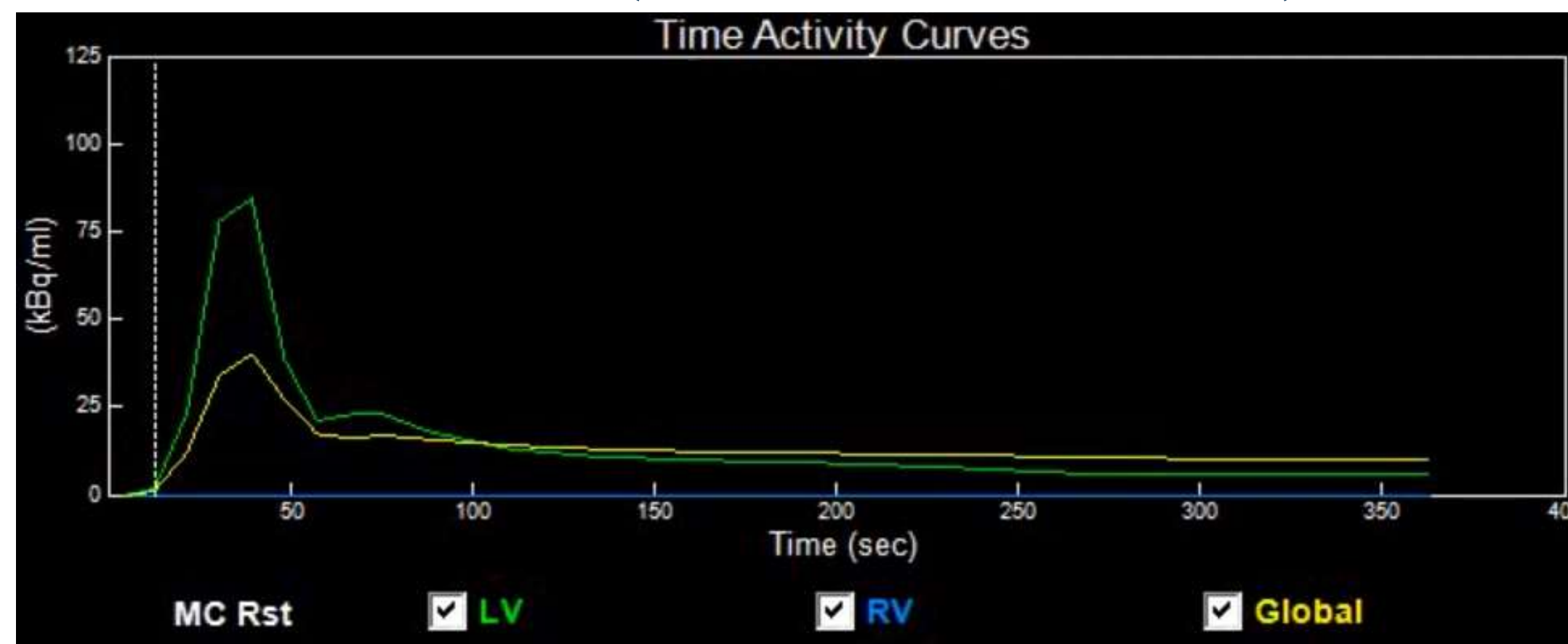


MPI WITH FULL-RING CZT SPECT

6-min 3D Dynamic SPECT/CT Acquisition (^{99m}Tc-Sestamibi)

- Rest study
- Bolus Injection (20 mCi)
- 67y Male
- BMI of 30.7

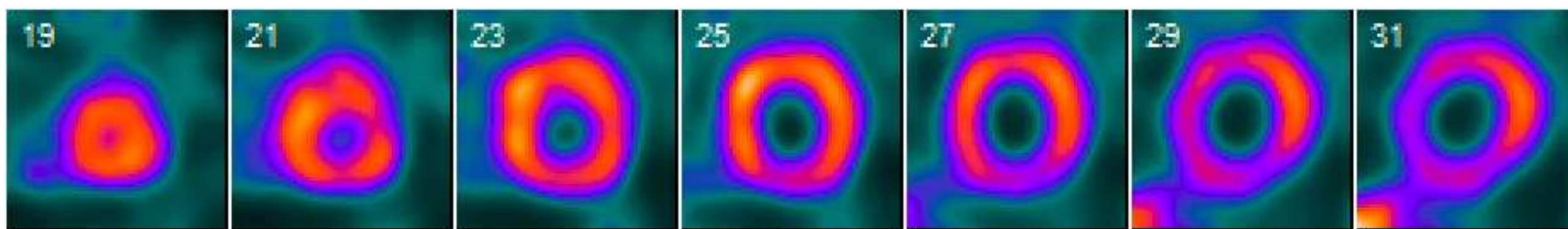
- Framing: 9x8 sec, 1x16 sec, 2x24 sec, 7x32 sec



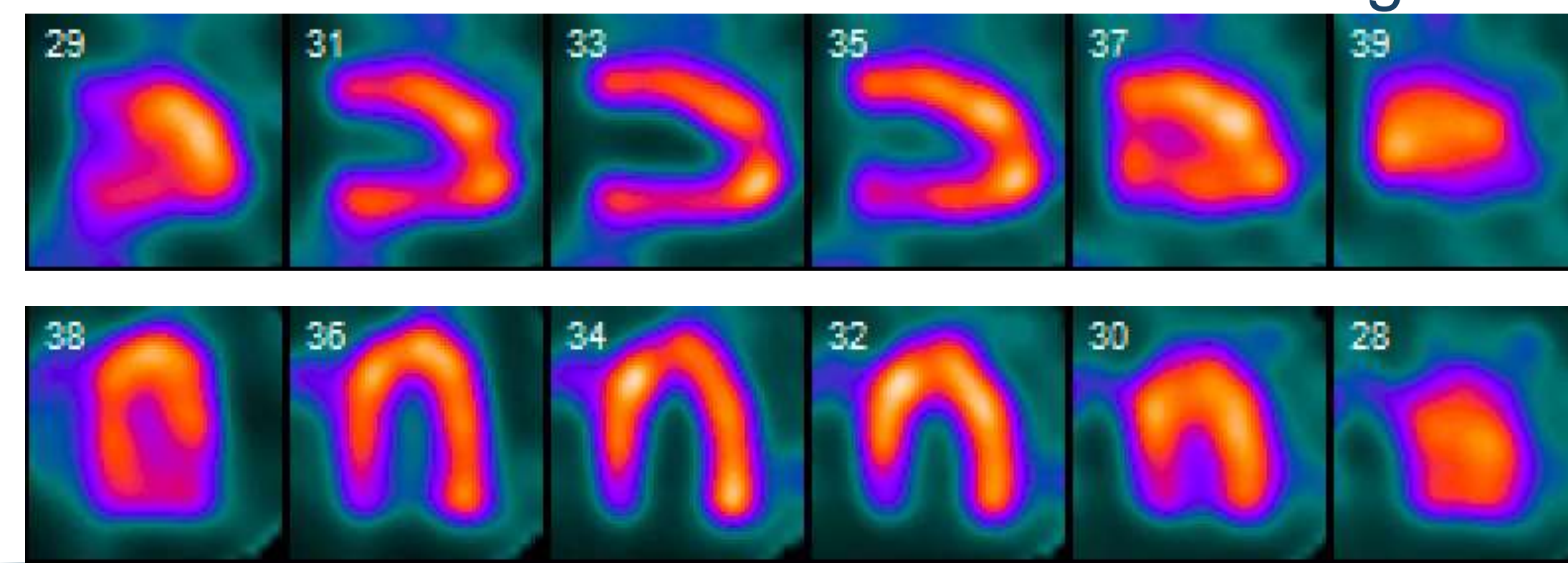
Net Retention Model

MBF (ml/min/g) ~ 0.62

9-min Gated SPECT/CT Acquisition

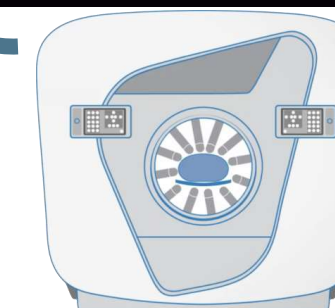


Short Axis



Horizontal Long Axis

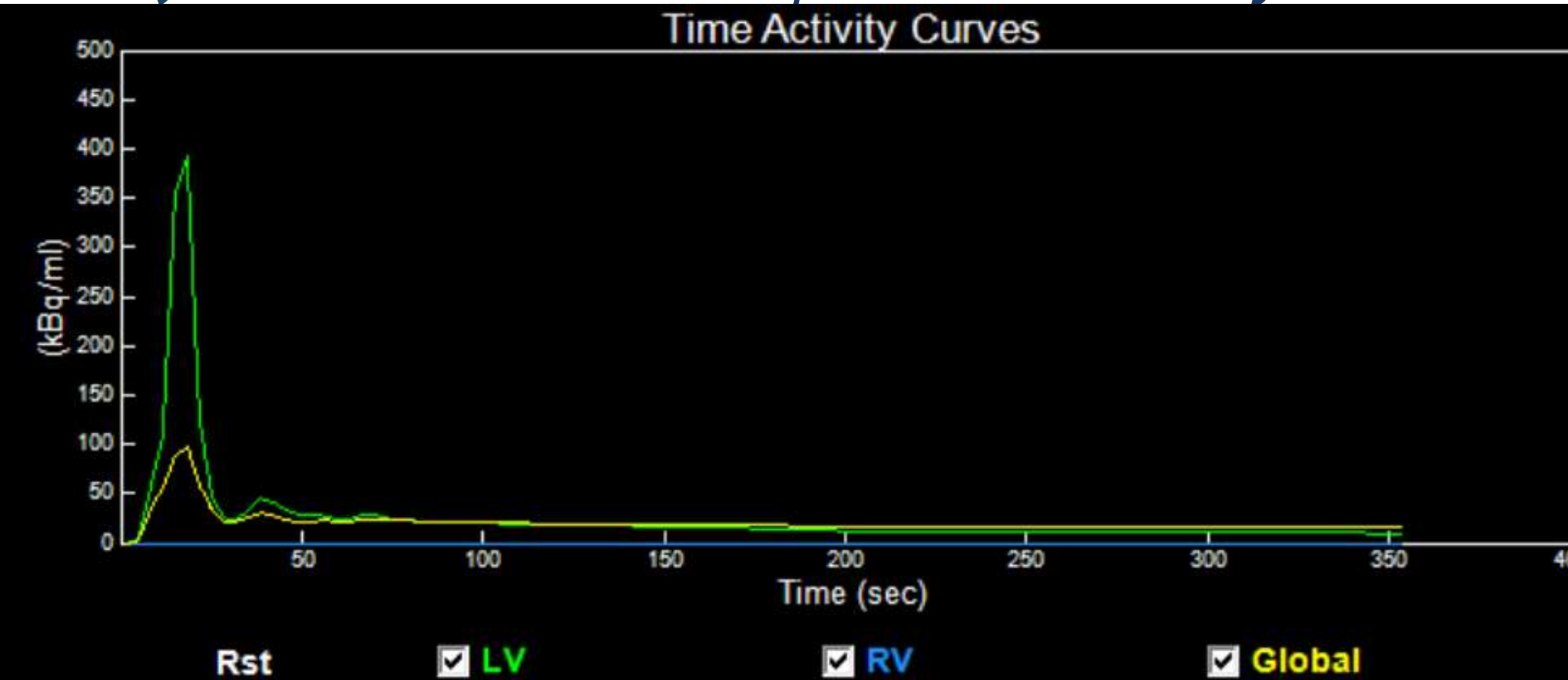
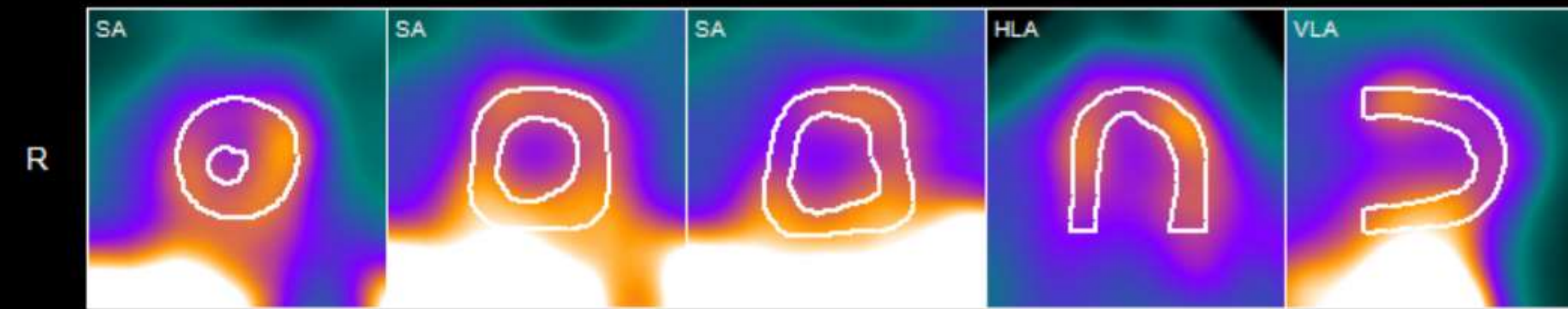
Vertical Long Axis



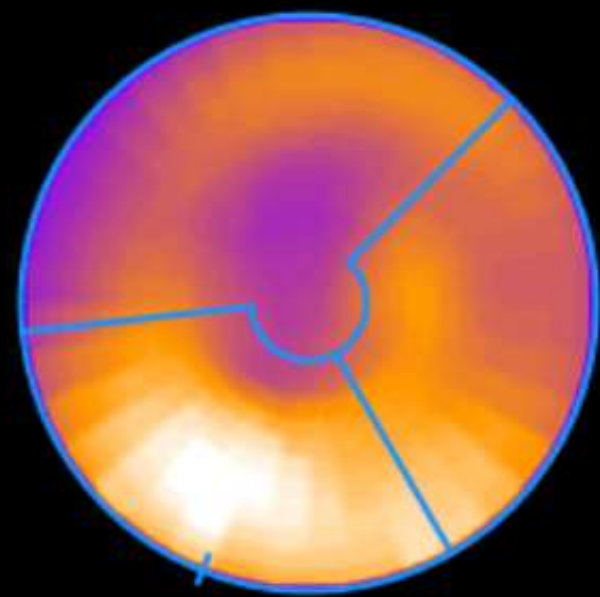


MPI WITH FULL-RING CZT SPECT

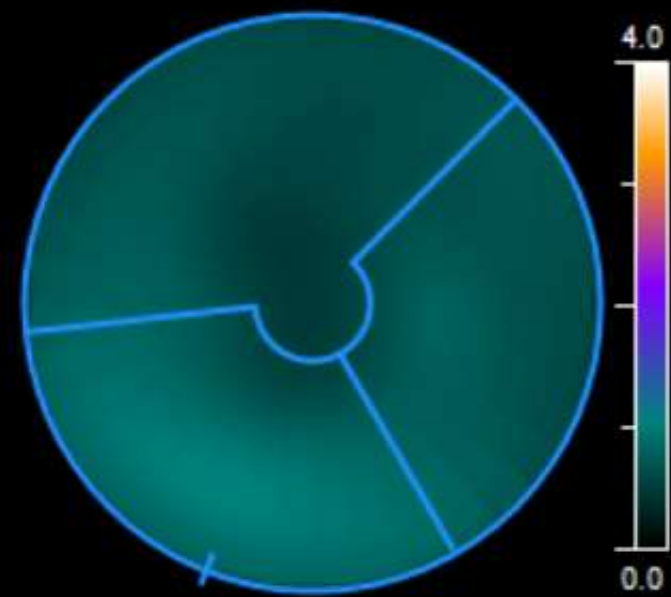
6-min 3D Dynamic SPECT/CT Acquisition (^{99m}Tc-Sestamibi) – Another Example Rest-Only



Perfusion-Normalized
RST1_DYN_2i16s-MBF_BP1_TA



Flow
RST1_DYN_2i16s-MBF_BP1_TA

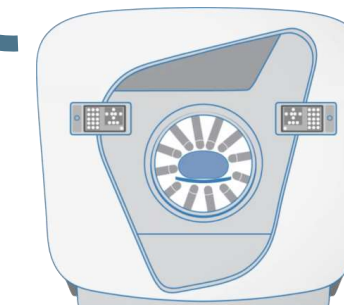


Flow (ml/min/g)
RST1_DYN_2i16s-MBF_BP1_TA

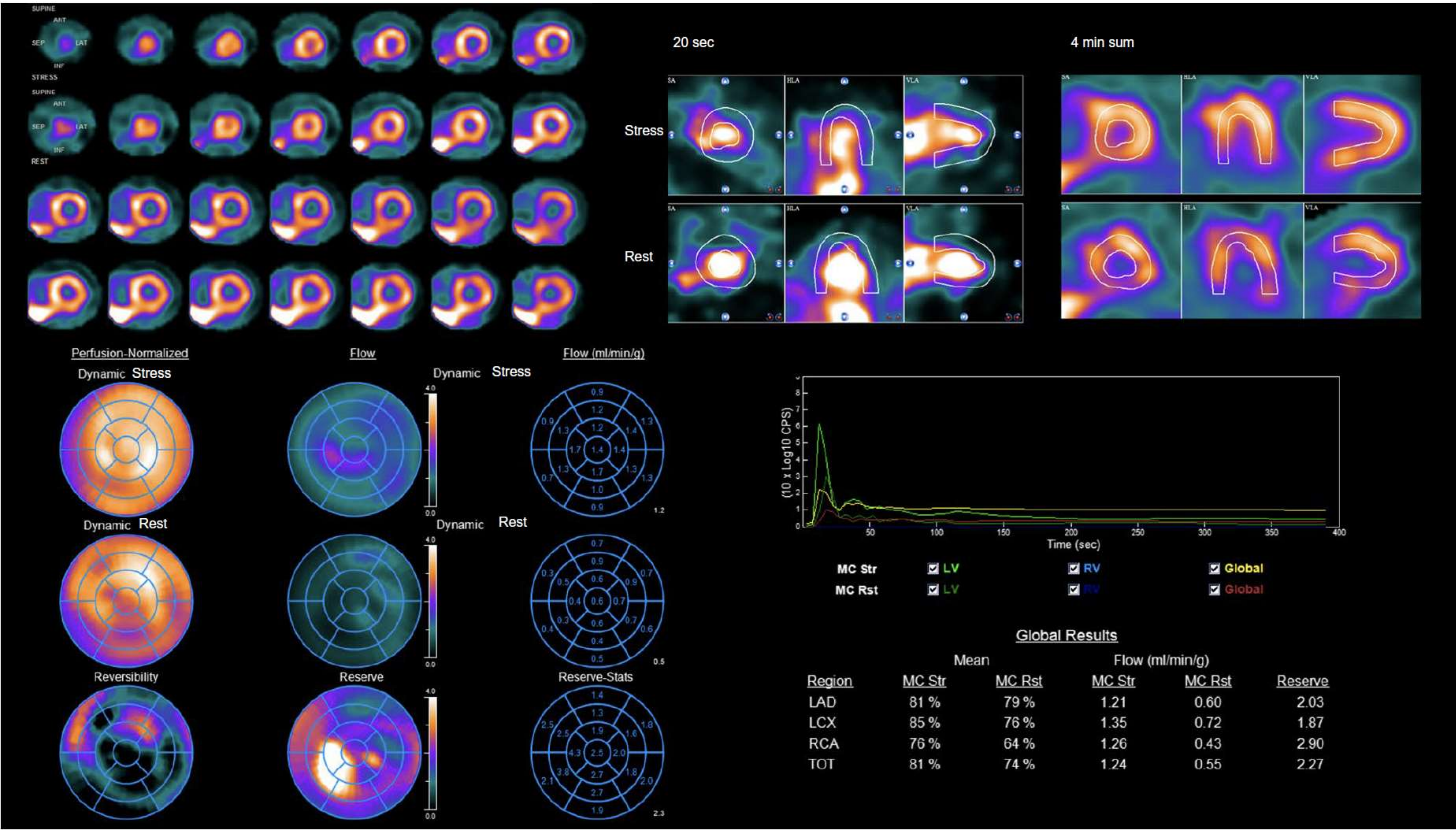


Algorithm (Rst): VERITON MIBI ROI NetRet Leppo AC
Algorithm (-): --

Region	Global Results				
	Mean		Flow (ml/min/g)		Ratio
	Rst	=	Rst	=	
LAD	70%	--	0.61	--	--
LCX	78%	--	0.69	--	--
RCA	86%	--	0.81	--	--
TOT	77%	--	0.68	--	--



7-min 3D Dynamic SPECT/CT Acquisition (^{99m}Tc-Sestamibi) – GE Starguide



Bailly, et al. (2023). Dynamic cardiac SPECT with flow measurement using 3D-ring CZT: when SPECT is inspired by PET. *EJNMMI*, 50(6), 1837-1839.



Cardiac Amyloidosis is a form of **progressive heart failure** (*protein misfolding disorder*)

Scintigraphy and SPECT/CT with Bone Avid Tracers (^{99m}Tc PYP/DPD),

❑ **>91% sensitive, 87% specific for cardiac ATTR [1]**

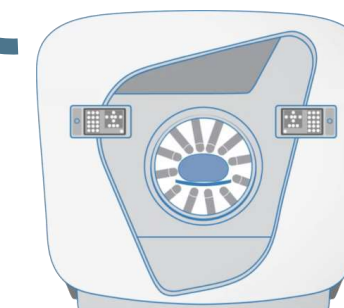
❑ **Standard of Care for Diagnosis [2]**

❑ **Potential Roles**

- **Prognosis**
- **Early diagnostic**
- **Assessment of disease progression**
- **Assessment of treatment response**

[1] Gillmore JD, et al. *Circulation* 2016.

[2] Dorbala, et al. *Circulation: Cardiovascular Imaging*, 2021.



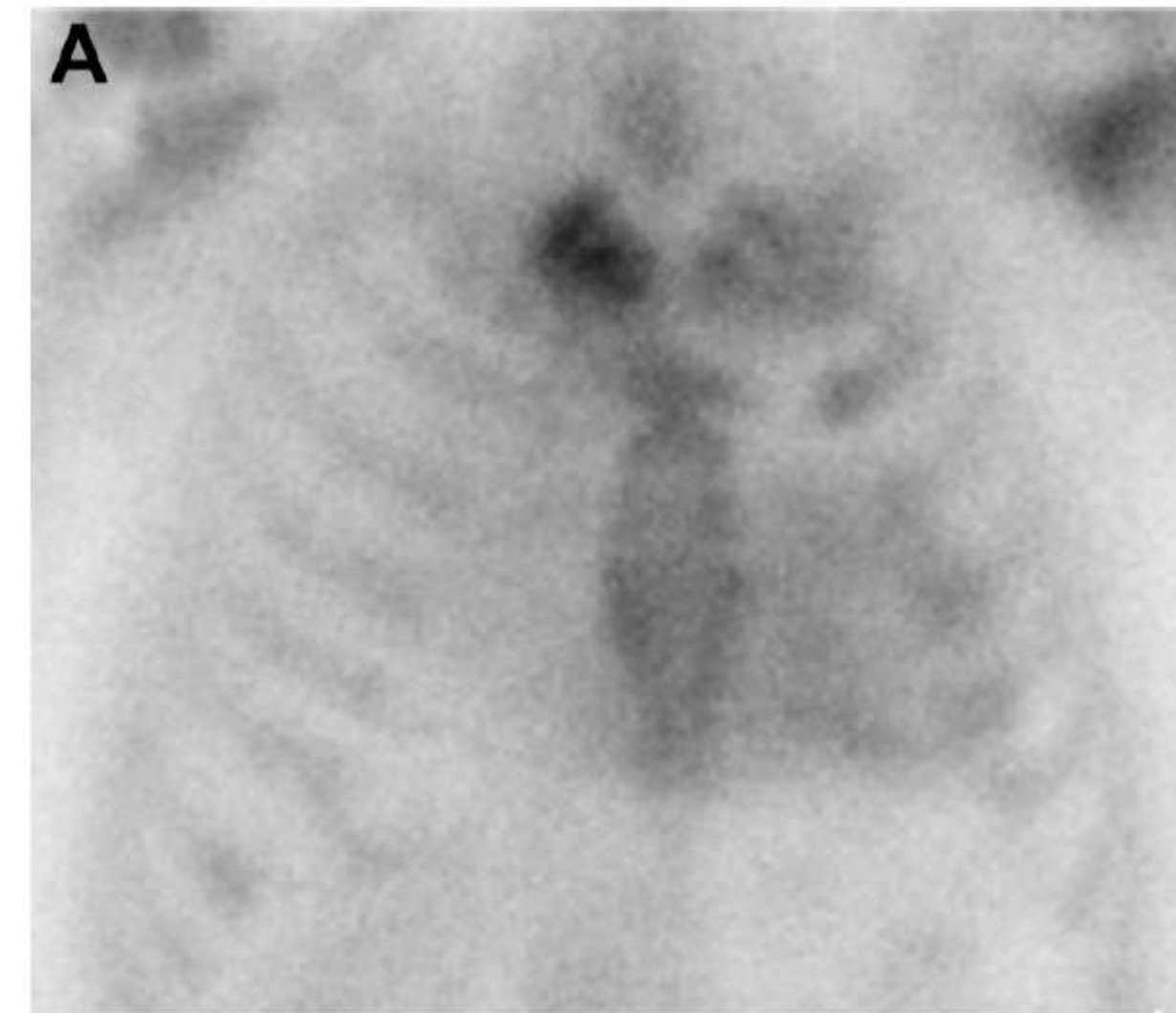
SPECT is recommended over scintigraphy [1,2],

- Delineate blood pool from myocardial activity

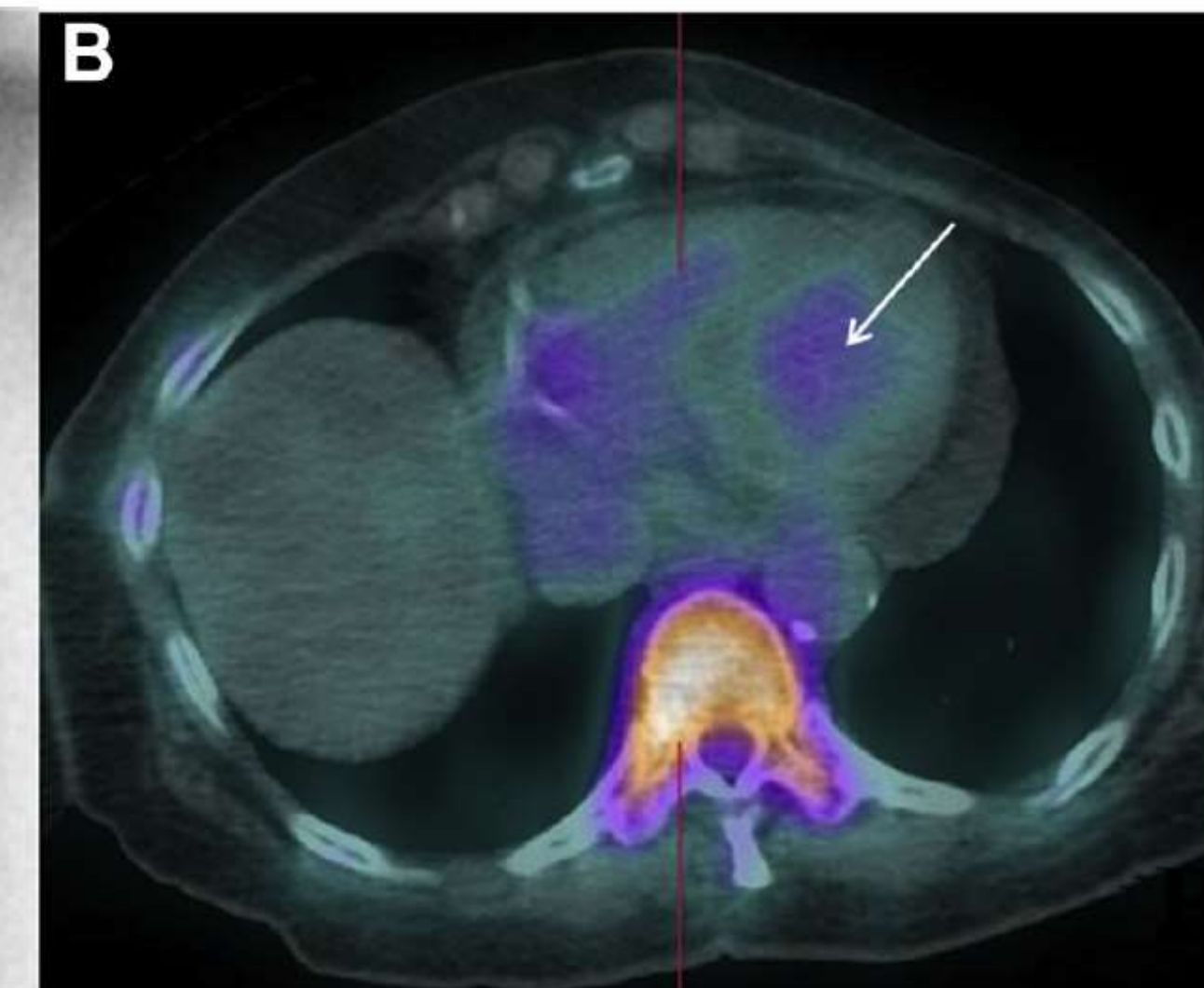
SPECT/CT is superior to SPECT [1,2]

- Attenuation and Scatter Correction
- Improved Quantitation
- Anatomic localization – particularly valuable for low-uptake scans and blood pool delineation

Scintigraphy
(*planar*)



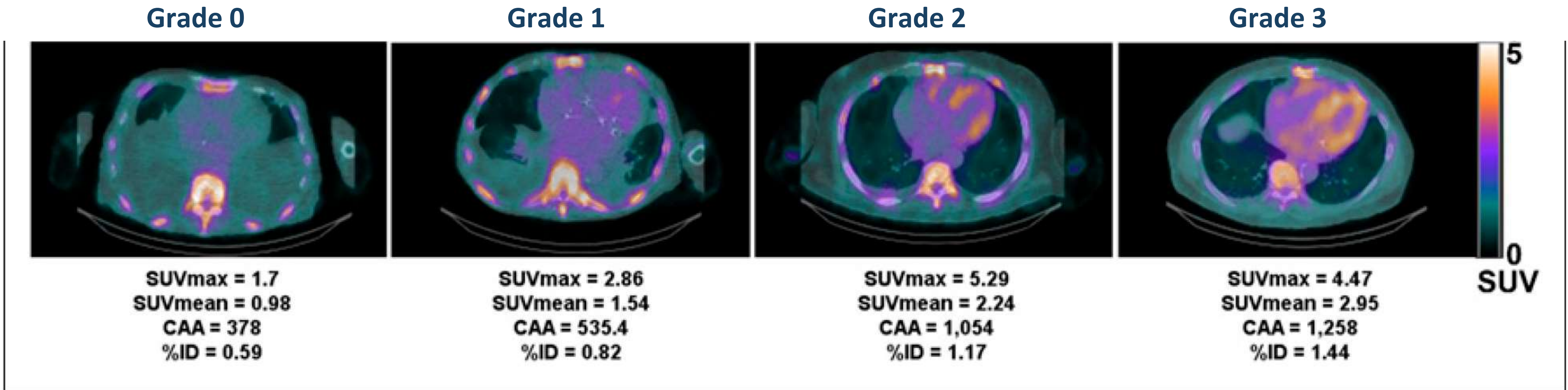
SPECT/CT



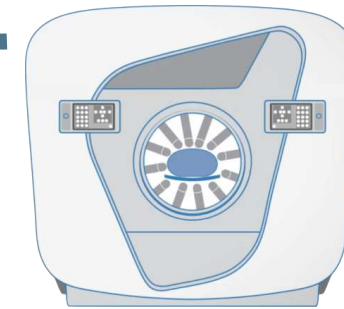
Adapted from [2] Hanna, et al. *Journal of the American College of Cardiology*, 2020.

[1] Dorbala, et al. *Circulation: Cardiovascular Imaging*, 2021.

Quantitative Full-Ring CZT SPECT/CT offers the potential for **early detection, risk stratification, and monitoring of response to therapy** in cardiac amyloidosis patients



Adapted from Dorbala, et al. *Journal of Nuclear Medicine*, 2021

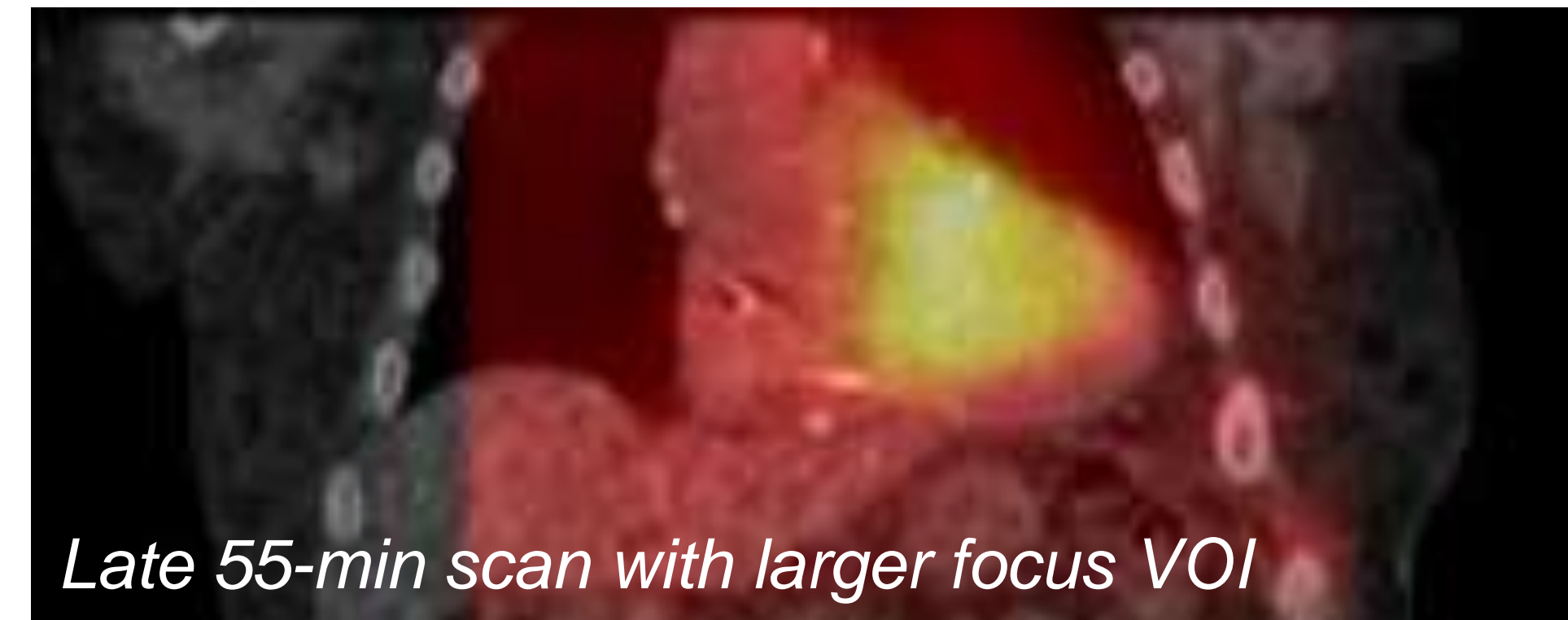
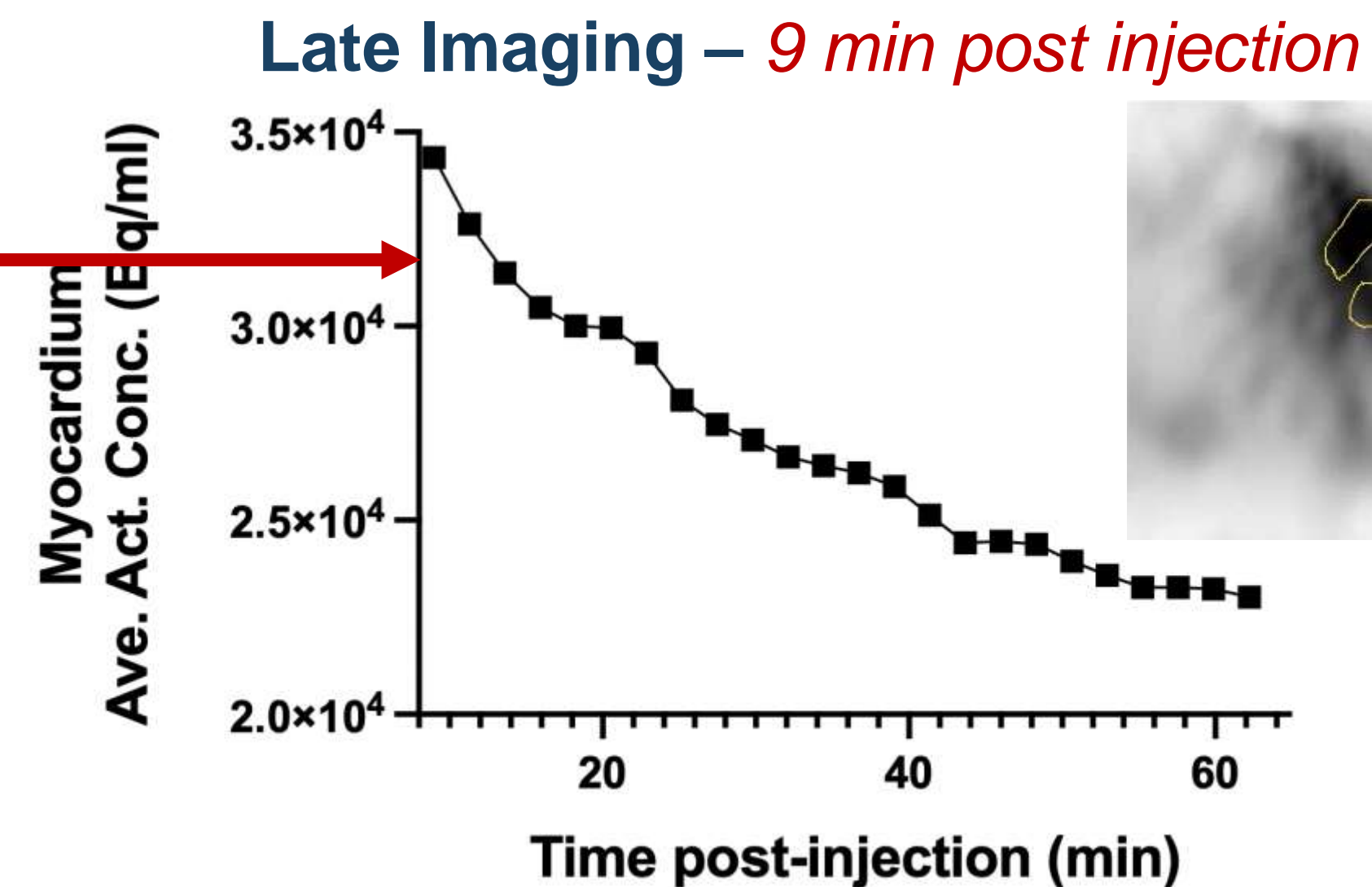
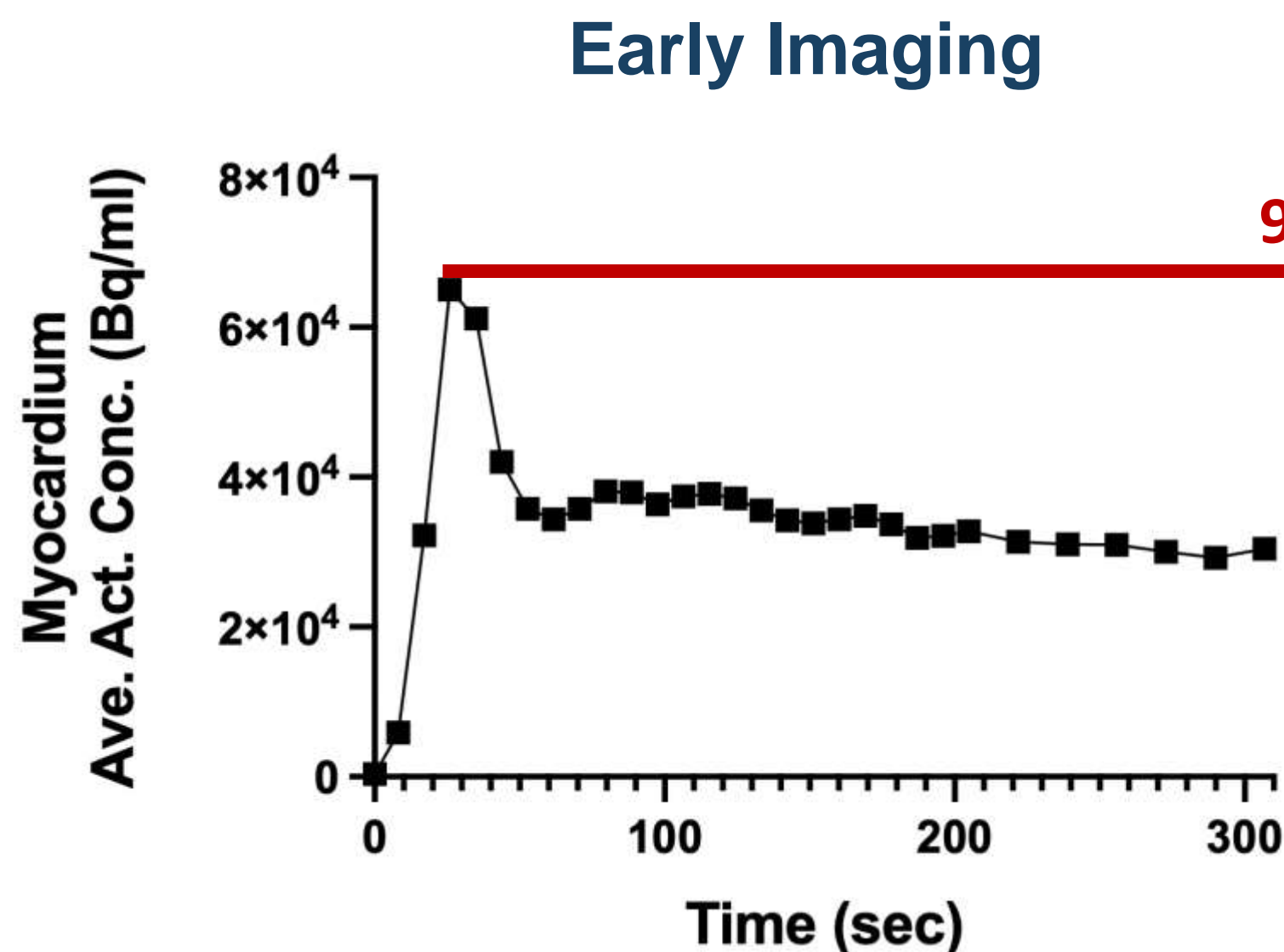


3D Dynamic Imaging enables,

- ❑ Segmentation of blood pool region from early frames to improve quantitation
- ❑ Determination of PYP/DPD myocardium uptake over time to reduce uptake time

Dynamic SPECT/CT

20 mCi (^{99m}Tc-PYP)
82y Male
Positive Case
BMI: 24.7

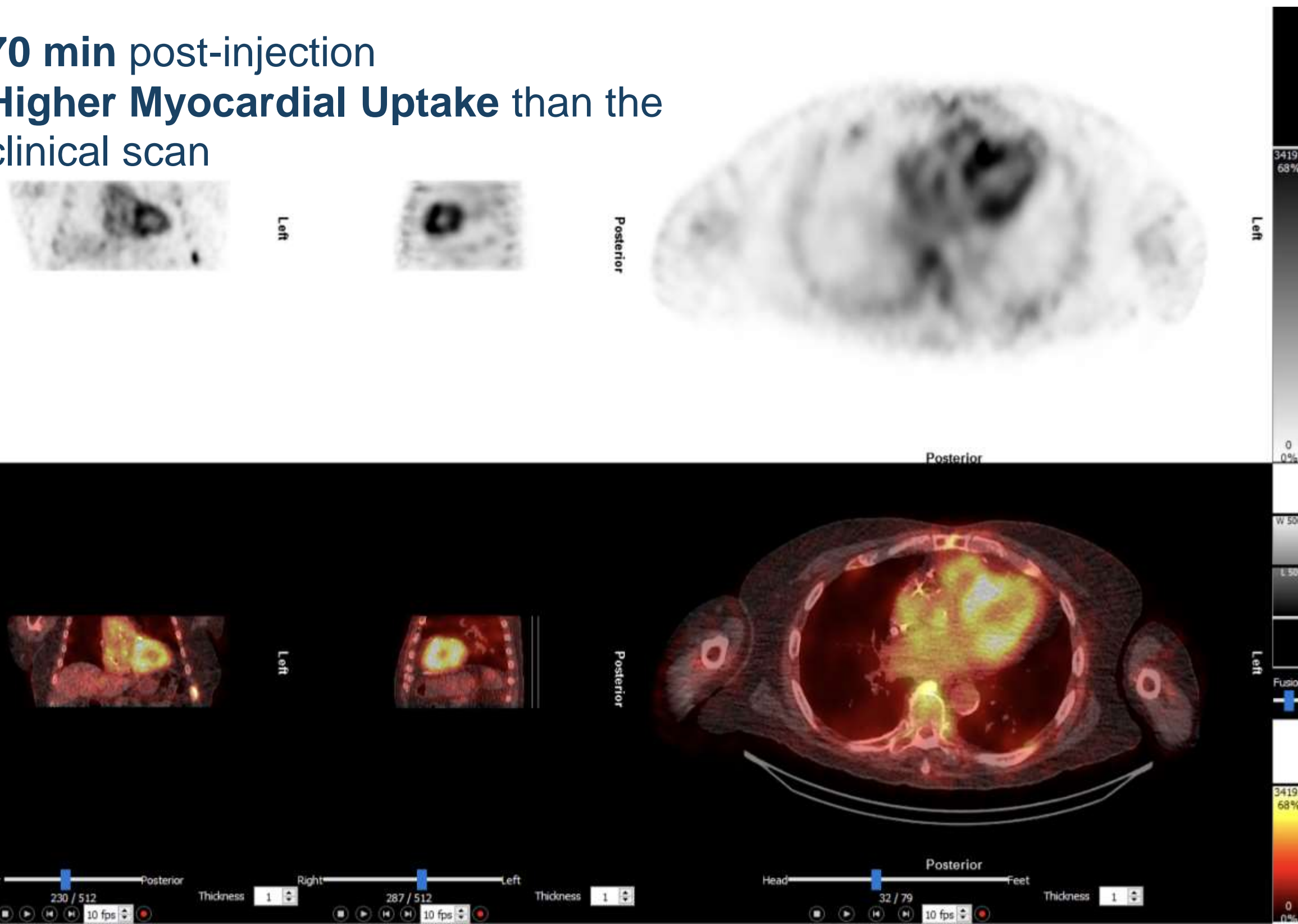


3D Dynamic Imaging enables,

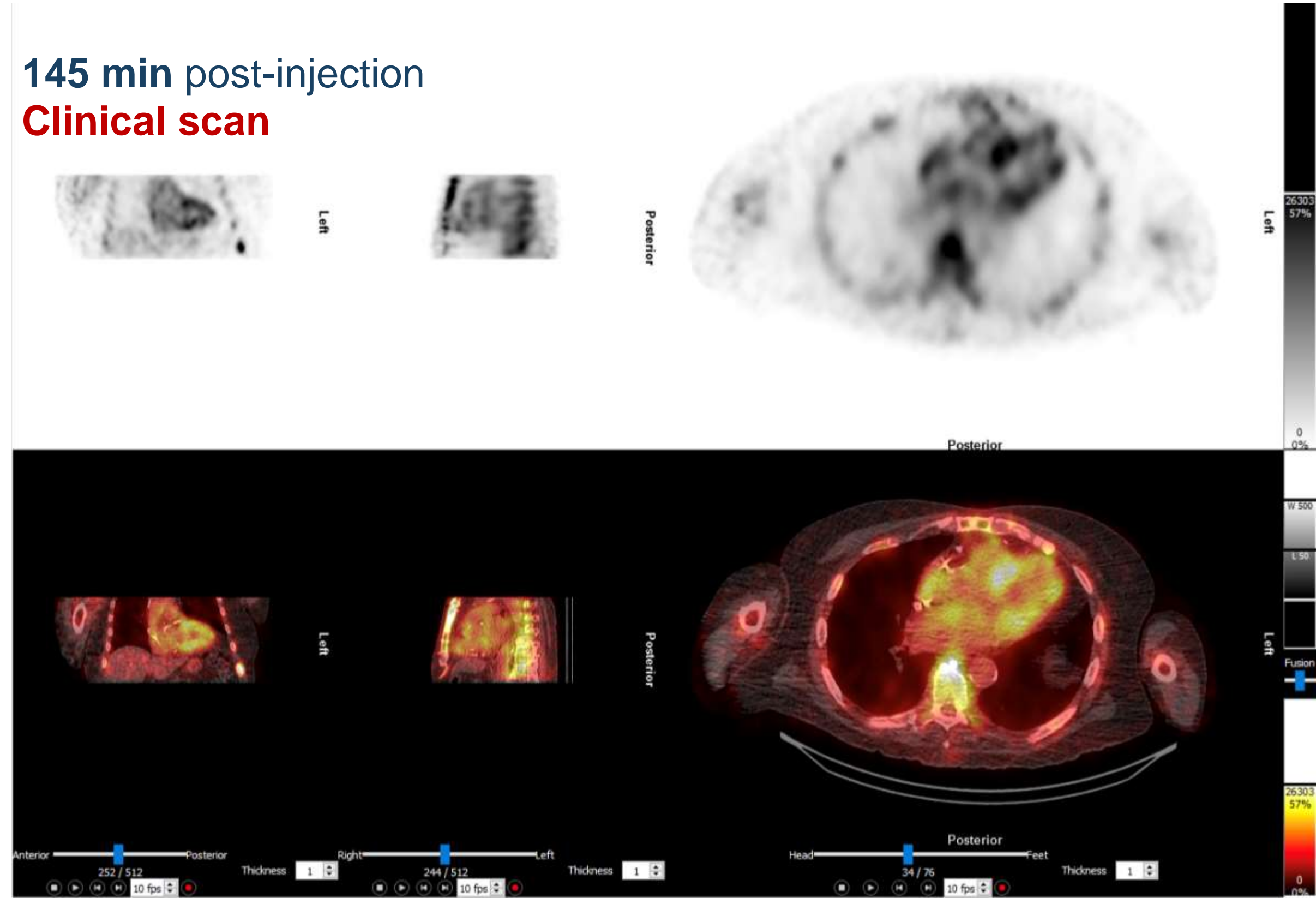
- ❑ Segmentation of blood pool region from early frames to improve quantitation
- ❑ Determination of PYP/DPD myocardium uptake over time to reduce uptake time

Cardiac Uptake already seen **early on**
Uptake time can very likely be **shortened**, resulting in **improved Clinical Care**

70 min post-injection
Higher Myocardial Uptake than the clinical scan



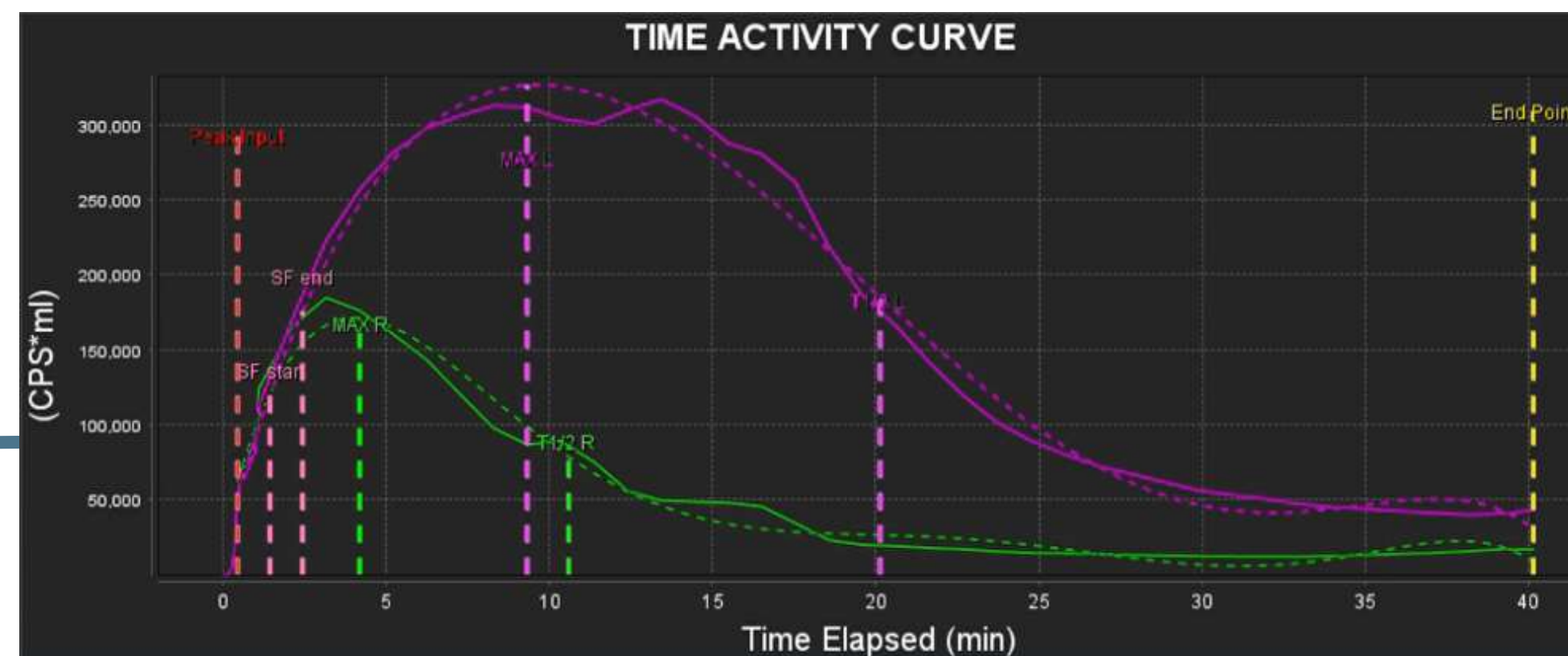
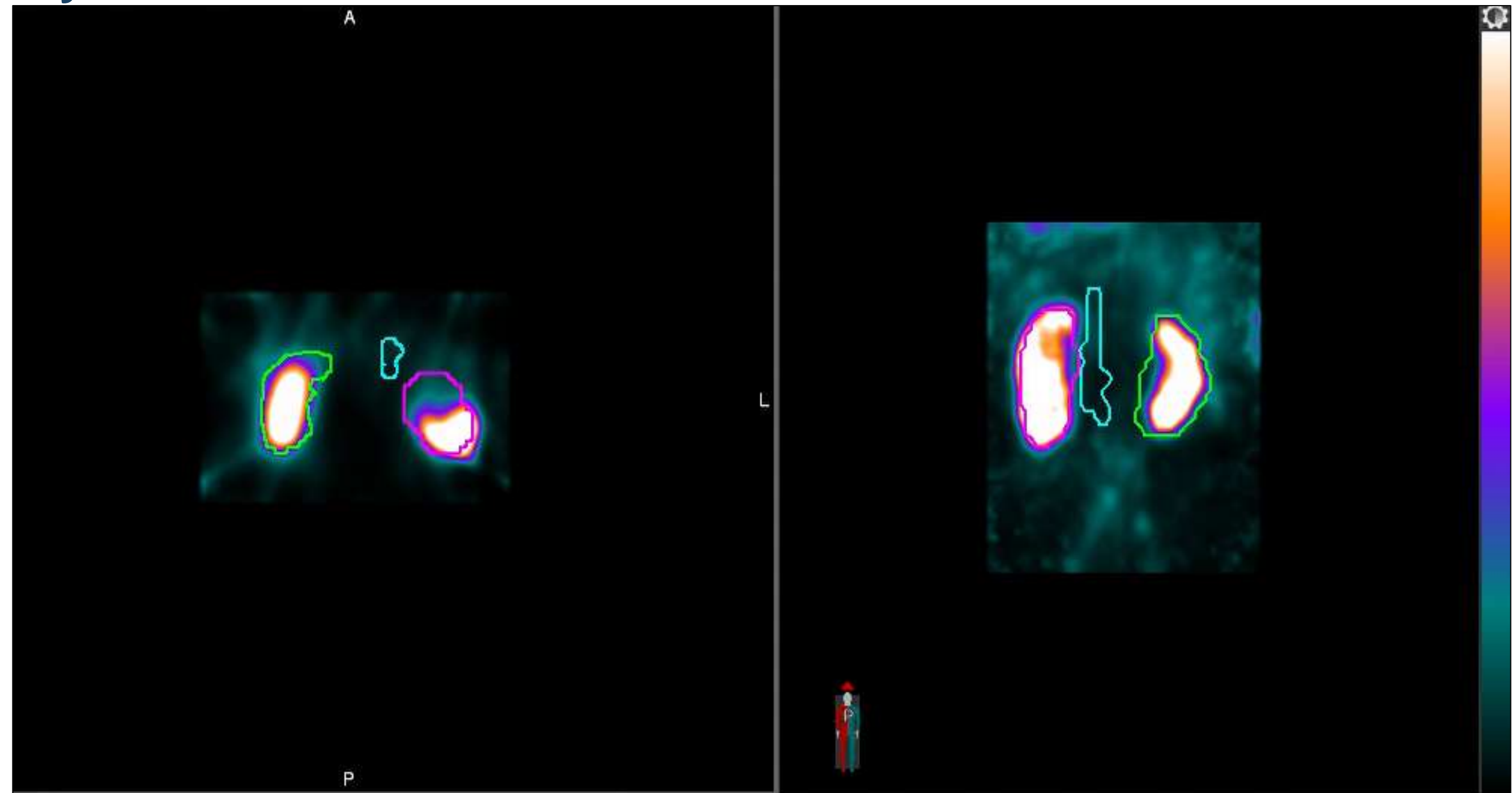
145 min post-injection
Clinical scan



Full-ring CZT SPECT/CT offers

- High sensitivity and improved spatial resolution that can enhance image quality and quantitation while reducing the acquisition time and/or injected dose
- CT-based anatomic information that can be used for ROI definition and to aid localization [1-3]
- Enhanced 3D dynamic capability

Dynamic SPECT/CT 40 min total scan time



84y Male; BMI: 24
^{99m}Tc-MAG-3: 10 mCi

Images courtesy of Royal United Hospitals Bath NHS Foundation Trust, Bath, UK

[1] Jacene, *et al.* Open Med Imaging 2008.
 [2] Arun, *et al.* Nucl. Med. Commun. 2013.
 [3] Eldredge, *et al.* Obesity Surgery. 2020.



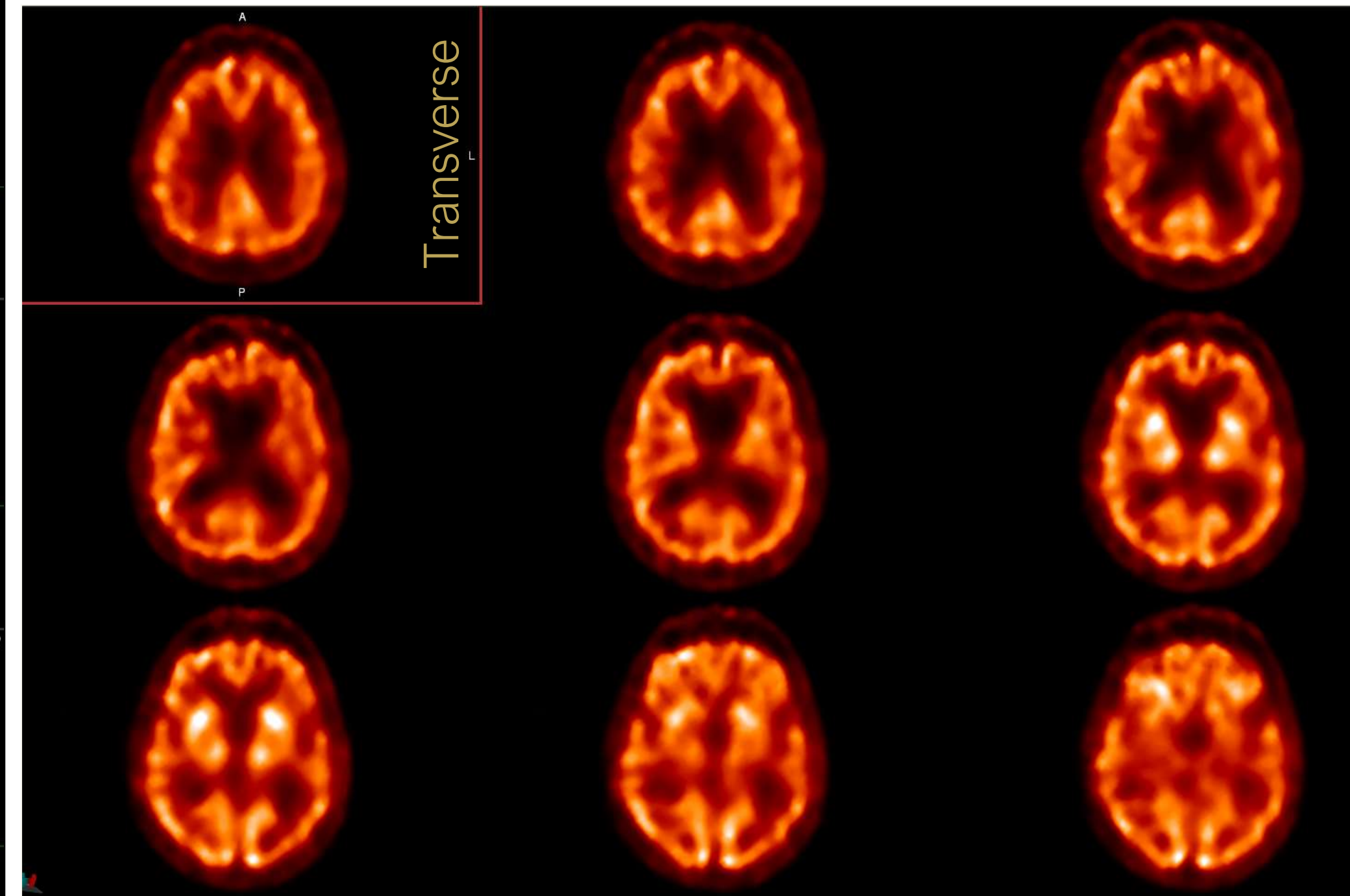
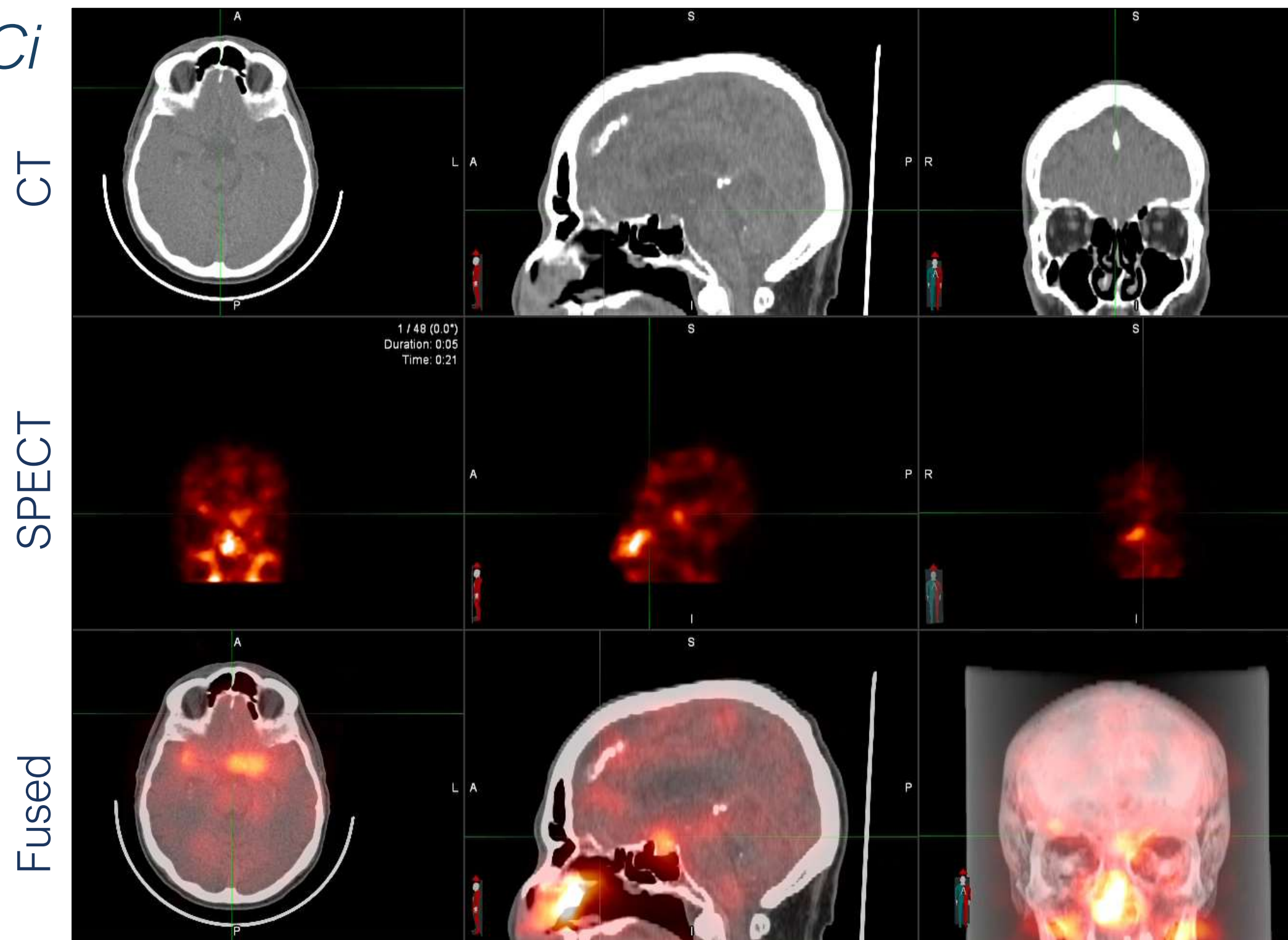
3D DYNAMIC BRAIN PERFUSION IMAGING

Dynamic SPECT/CT Acquisition (^{99m}Tc-ECD)

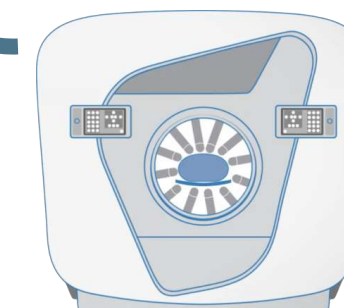
- 79y Male
- BMI: 21
- Dose: 17.9 mCi

Dynamic SPECT/CT 5 sec/frame – 15 min

Delayed Static SPECT/CT – 15 min



Images Courtesy of Centre Hospitalier Régional Universitaire (CHRU) de Nancy, Nancy, France



Full-ring CZT SPECT/CT

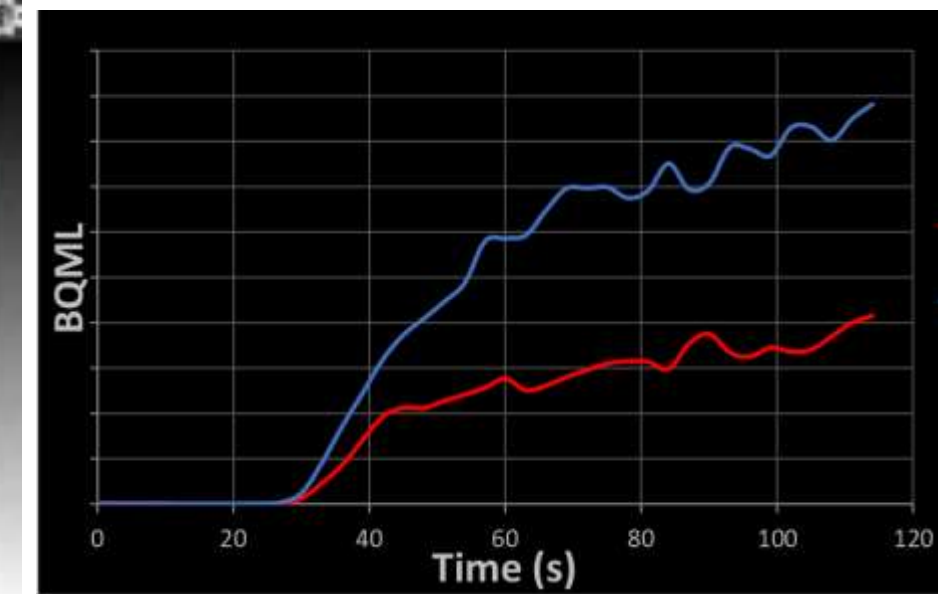
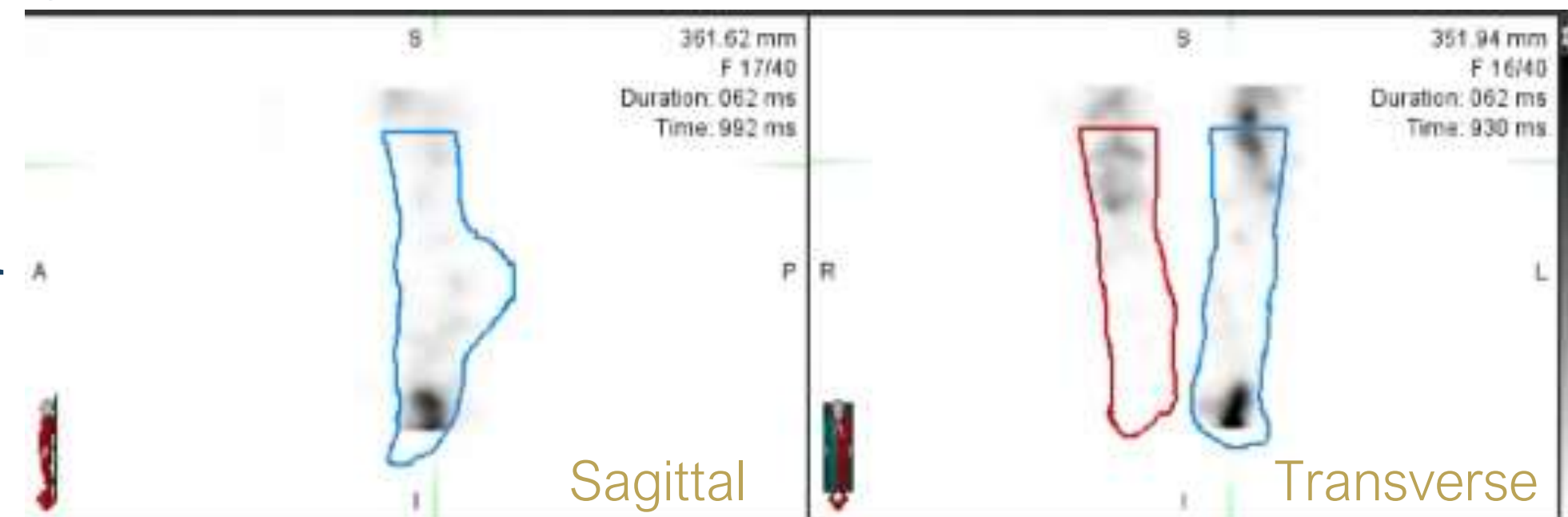
Can offer **CT-based anatomic information** that can be used for ROI definition compared to **SPECT-only** and **planar**

Provides **3D dynamic capability** that can enhance **multi-phase bone imaging** compared to **conventional SPECT/CT** (*limited to 2D Dyn Imaging*)

Three Phase Bone SPECT/CT

- 45y Male
- BMI: 26.6
- ^{99m}Tc-HMDP 12.1 mCi

Dynamic SPECT/CT 2 min (40 frames of 3 sec)



Flow phase (I)



Dynamic SPECT/CT
2 min

BP phase (II)



Static SPECT/CT
5 min

Delayed phase (III)



Static SPECT/CT
5 min

Mairal *et al.* . Eur J Nucl Med Mol Imaging.

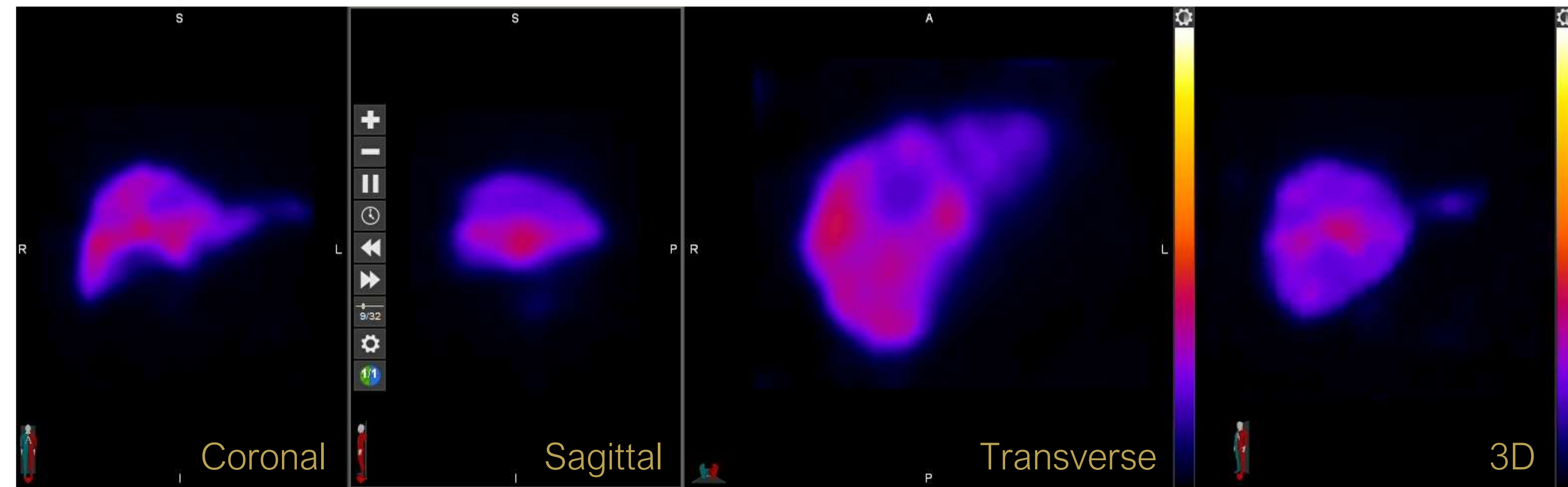
Full-ring CZT SPECT/CT offers

High sensitivity and improved spatial resolution that can enhance image quality and quantitation while reducing the acquisition time and/or injected dose

CT-based anatomic information that can be used for ROI definition and to aid localization [1-3]

Enhanced 3D dynamic capability

Dynamic SPECT 30 min (360 frames of 5 sec)



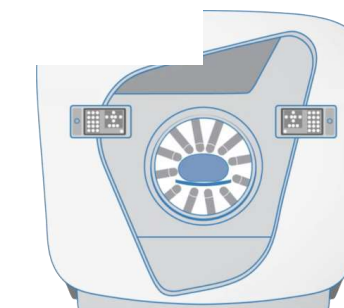
Planar CCK delayed at 60 min – 25% Ejection fraction

Dynamic SPECT-only followed by planar CCK

- 26y Female; BMI: 41.5
- ^{99m}Tc-diisopropyl IDA: 5 mCi



[1] Jacene, *et al.* Open Med Imaging 2008.
 [2] Arun, *et al.* Nucl. Med. Commun. 2013.
 [3] Eldredge, *et al.* Obesity Surgery. 2020.



Major Liver Resection can be performed with **limited morbidity and mortality** when sufficient remnant liver function remains to avoid **Post-Hepatectomy Liver Failure (PHLF)**.

Pre-operative assessment of the future remnant liver function **is essential** [1,2]

❑ **Current Standard – Indirect Imaging Approach**

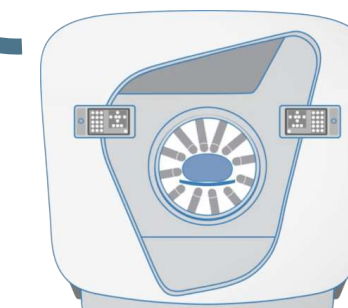
1. Dynamic Planar Imaging to assess total liver function (6 min)
2. Fast static SPECT/CT at liver maximum uptake to segment remnant/resected liver regions (10 min)

❑ **Role for Full-Ring CZT SPECT = 3D Dynamic Imaging**

- **Facilitate** the procedure - *single dynamic SPECT only*
- **Direct** liver remnant function **assessment**
- **Enhance** quantitative accuracy - *no activity superimposition*

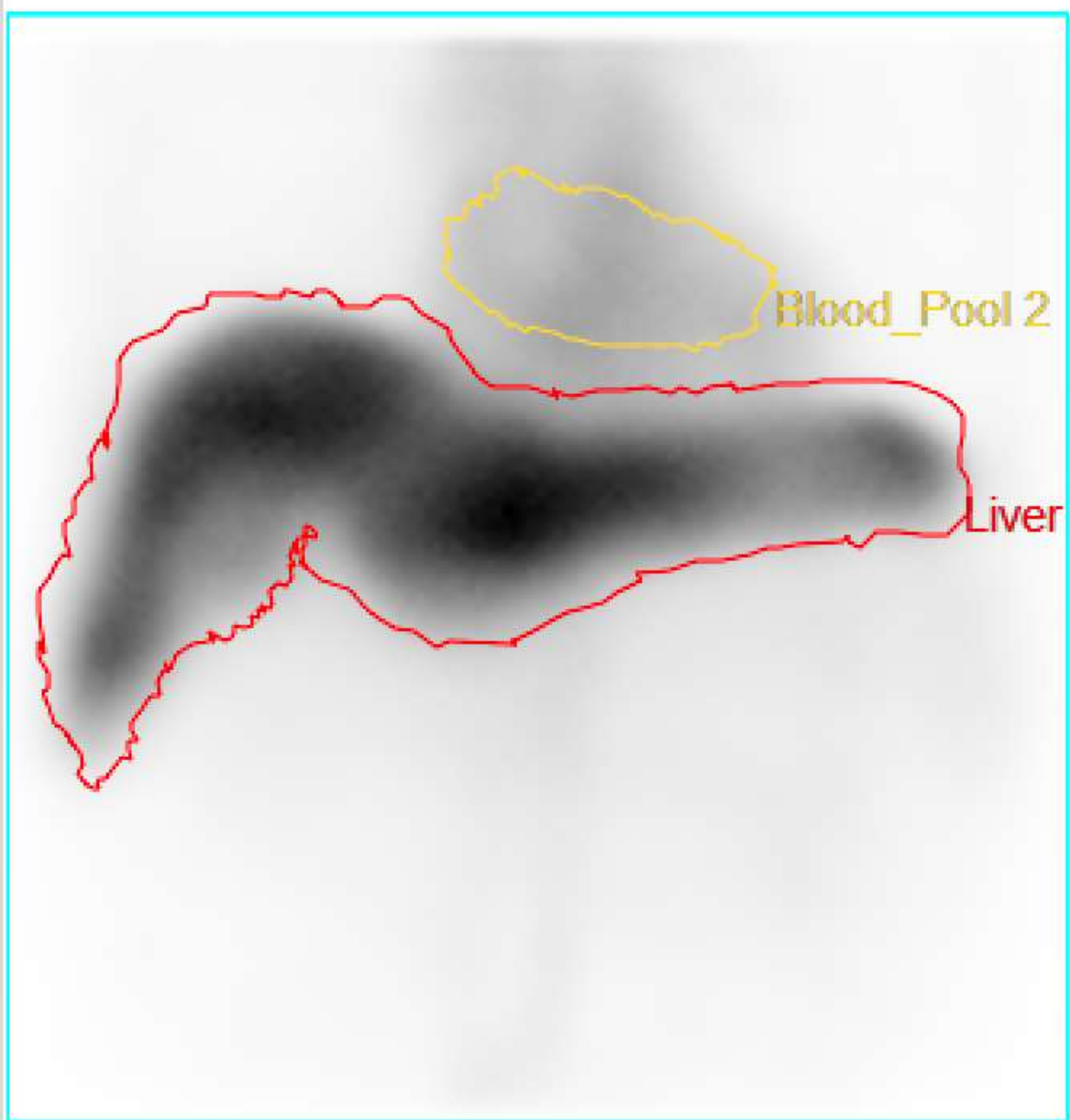
[1] Rassam, *et al.* Nuc. Med. Commun. 2019.

[2] Serenari, *et al.* Ann. Surg. 2018



Total Liver Function

- **^{99m}Tc-Mebrofenin (6.5 mCi)**
- **Dynamic Planar Imaging (10 sec/frame)**
- **Geometric mean from ANT and POST views to correct for source non-uniformities.**
- **ROIs for Liver, Blood Pool, and Total Body/FOV**

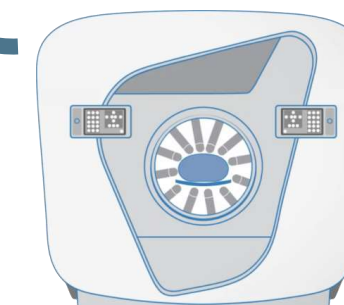
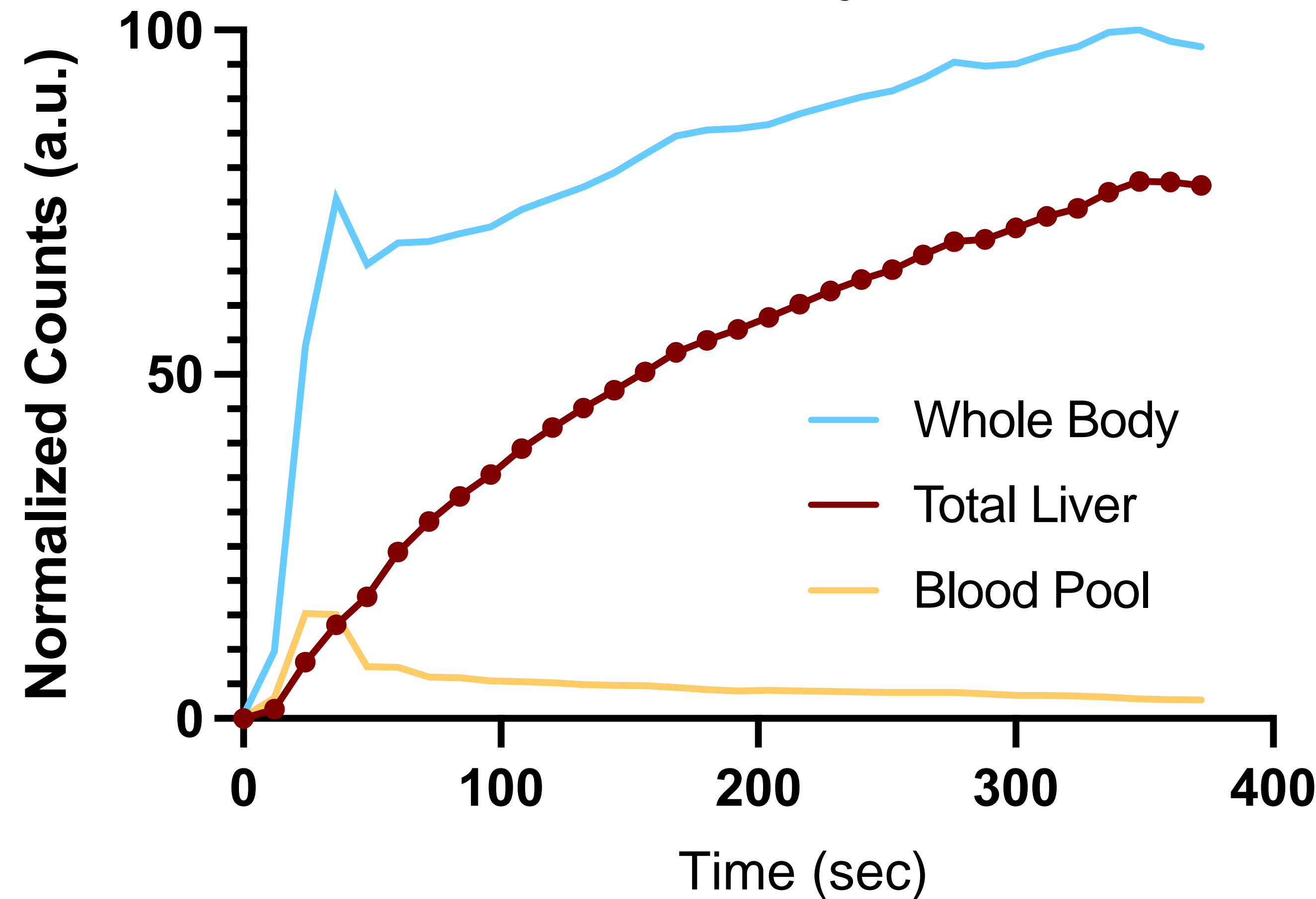


Dynamic Planar (2D)

6sec_BP1_MVP_Planar:D1
:20:59



Time Activity Curves

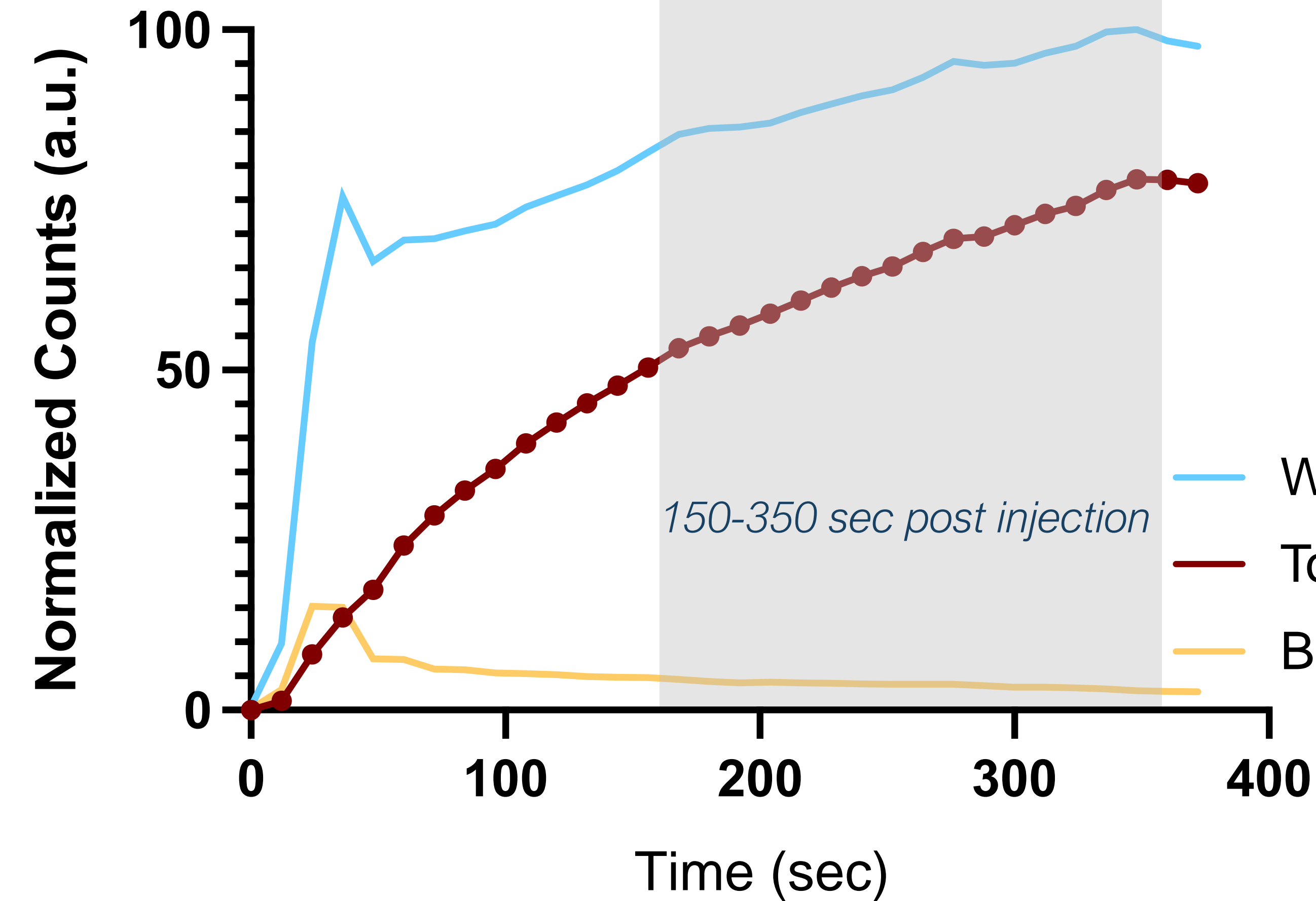




Total Liver Function Estimated from 2D Dynamic Imaging

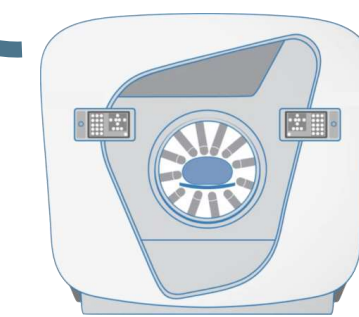
Time Activity Curves

Total Liver Uptake



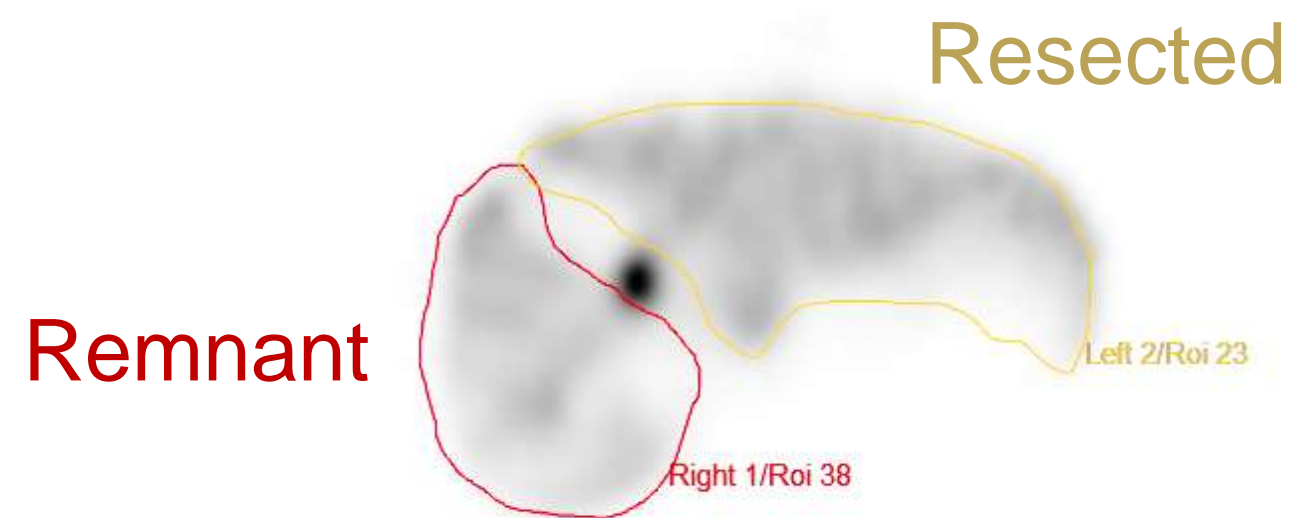
$$TL - U = 100 \times \frac{(AUC_{Liver} - AUC_{BP})}{AUC_{WB}}$$
$$TL - U = 66.8\%$$

- Whole Body
- Total Liver
- Blood Pool



Remnant/Resected Segmentation - % Counts from Static SPECT/CT

Transaxial SPECT

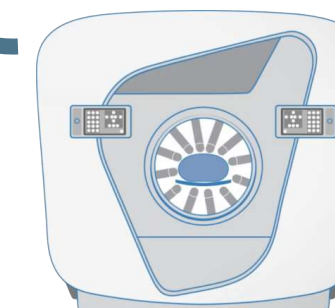
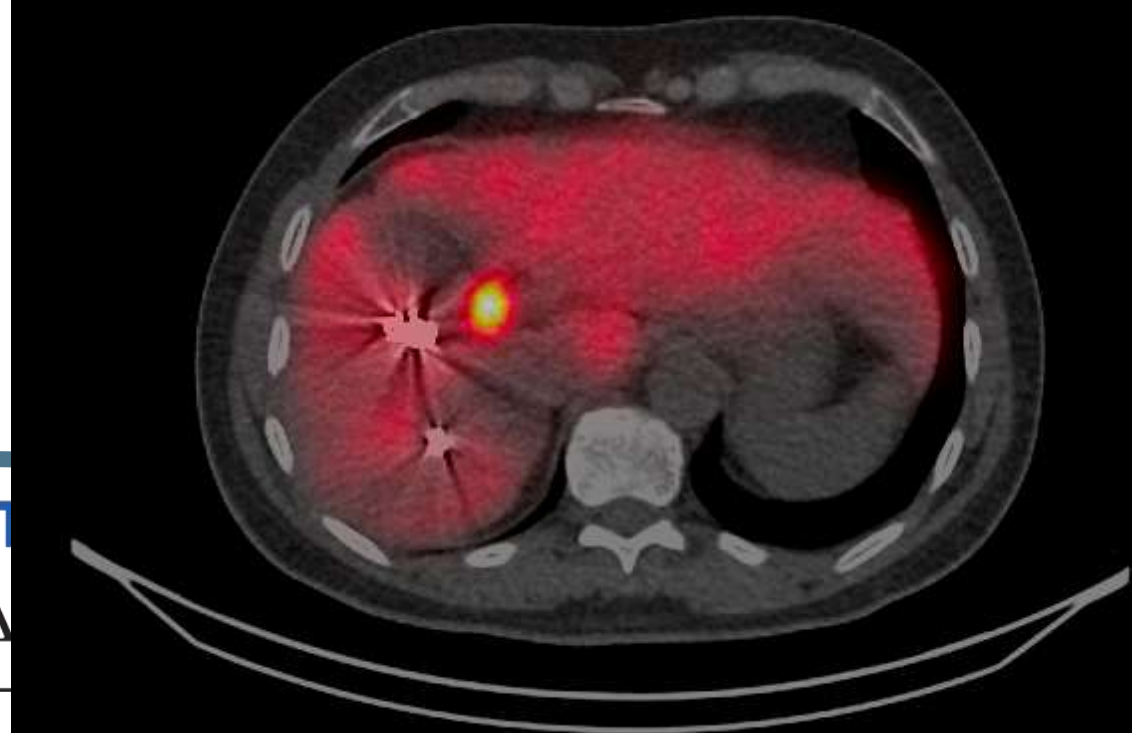


Future Remnant volume represents **52.7%** of liver function (counts)

3D View



Transaxial CT





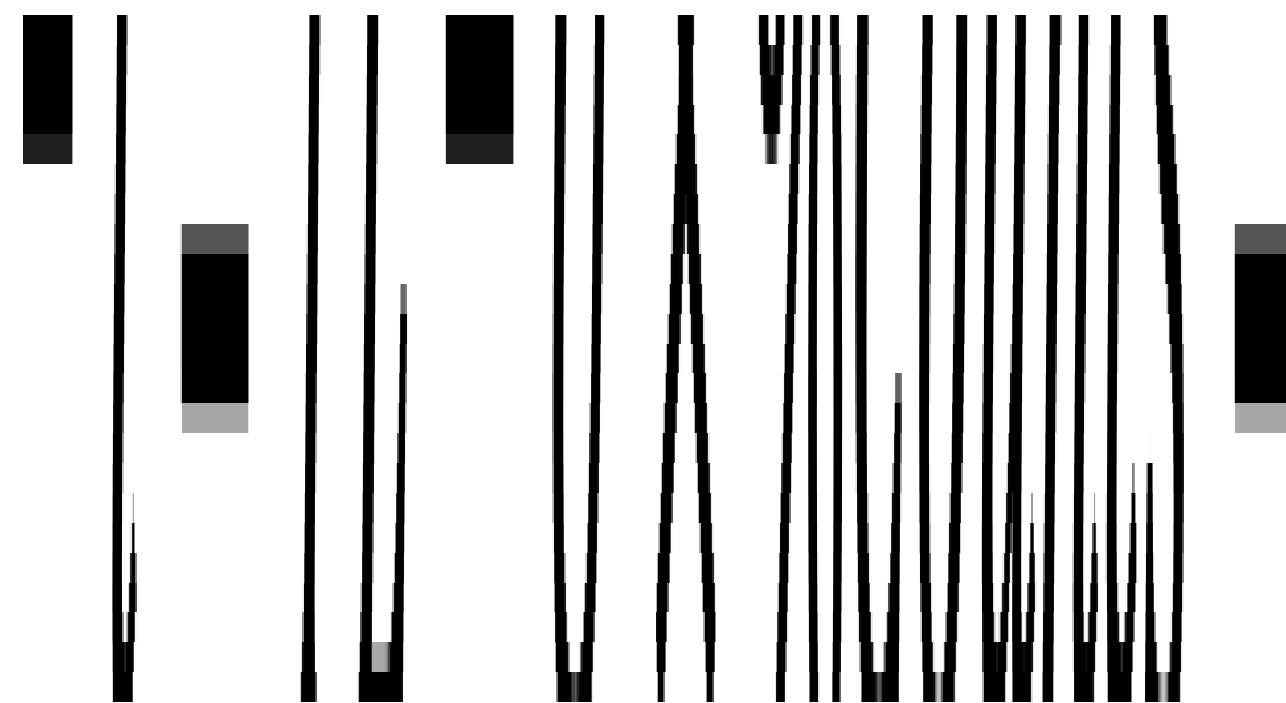
HIBA-*i* quantitation

Dynamic Planar



$$TL - U = 66.8\%$$

Static SPECT/CT



Future Remnant volume represents **52.7%** of liver function (counts)

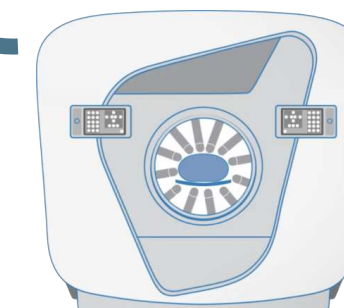
HIBA-*index*

used to predict *Post-Hepatectomy Liver Failure (PHLF)*

$$HIBA - i = TL - U \times \%Counts = 35.2\%$$

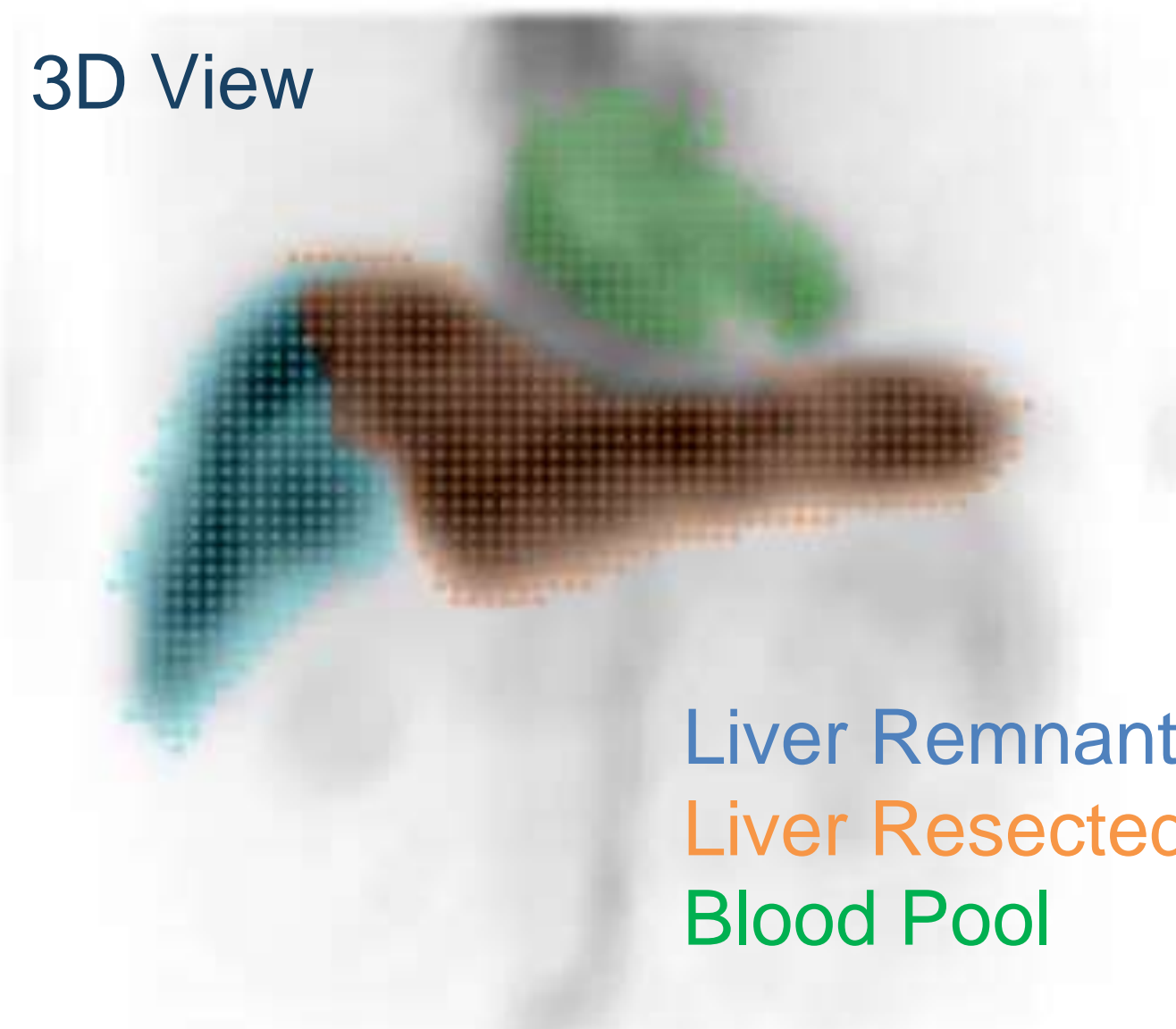
Compared against population-based cut-off values to determine risk of PHLF ($\geq 15\%$) [1,2]

- [1] Rassam, *et al.* Nuc. Med. Commun. 2019
- [2] Serenari, *et al.* Ann. Surg. 2018

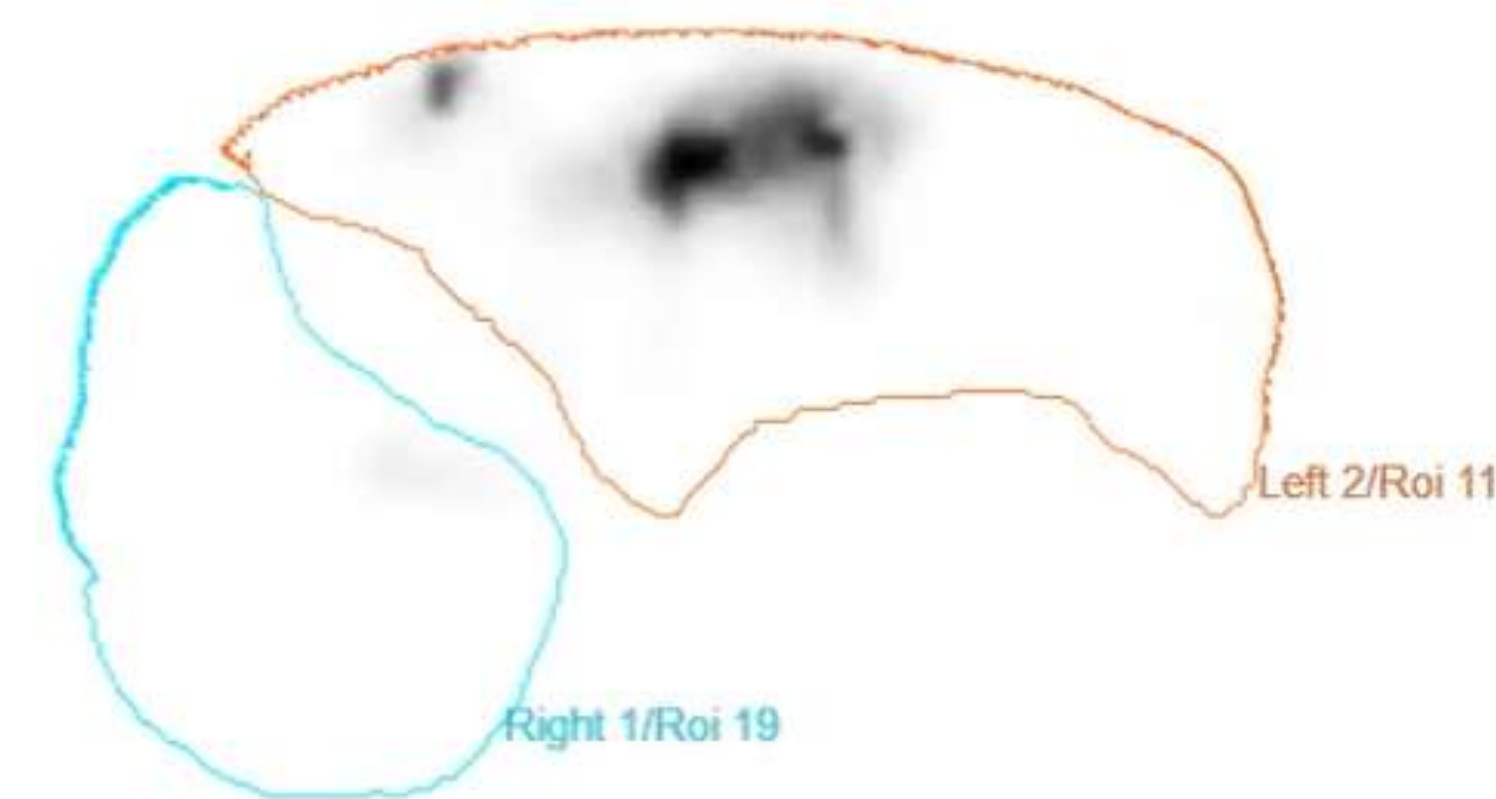


Full-ring CZT Hepatobiliary SPECT/CT,

- Enables 3D dynamic quantitative imaging
- Remnant liver function can be **directly** evaluated from 3D dynamic SPECT/CT
- More **accurate liver/remnant function estimation** since no superimposition of activity (*tends to be overestimated with planar imaging*)



Transaxial



Coronal

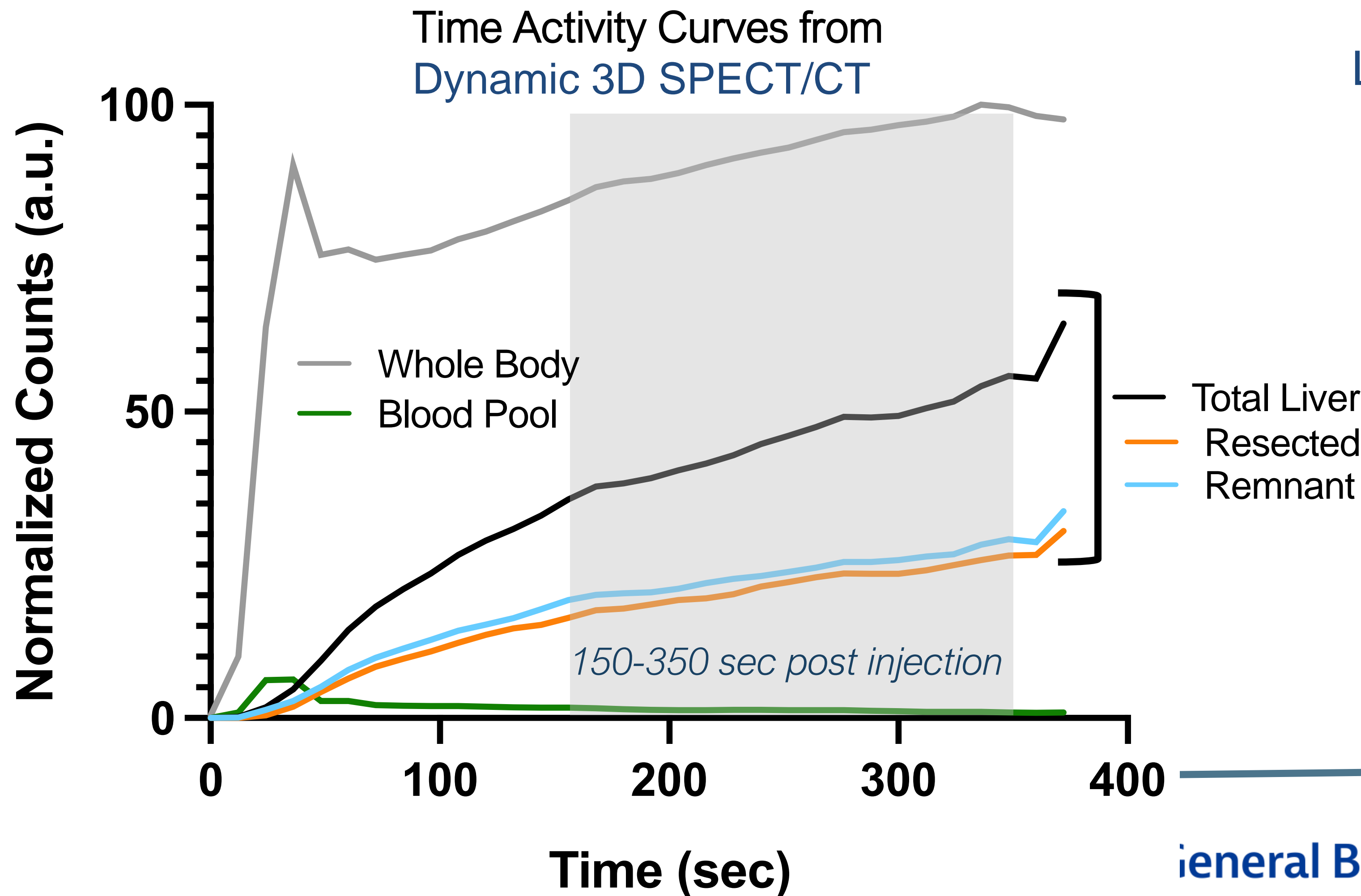


Sagittal





HIBA-i from 3D dynamic SPECT/CT



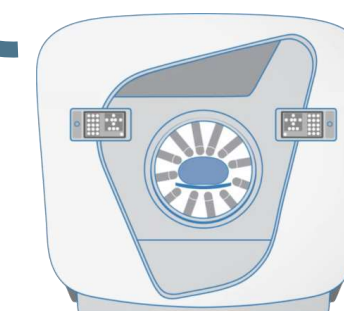
Liver Remnant Tracer Uptake

$$L - U_{Remnant} = HIBA - i = 24.9\%$$

$$L - U_{Resected} = 22.6\%$$

$$TL - U = 47.5\%$$

Will likely need establishment of new population-based cut-off values to determine PHLF

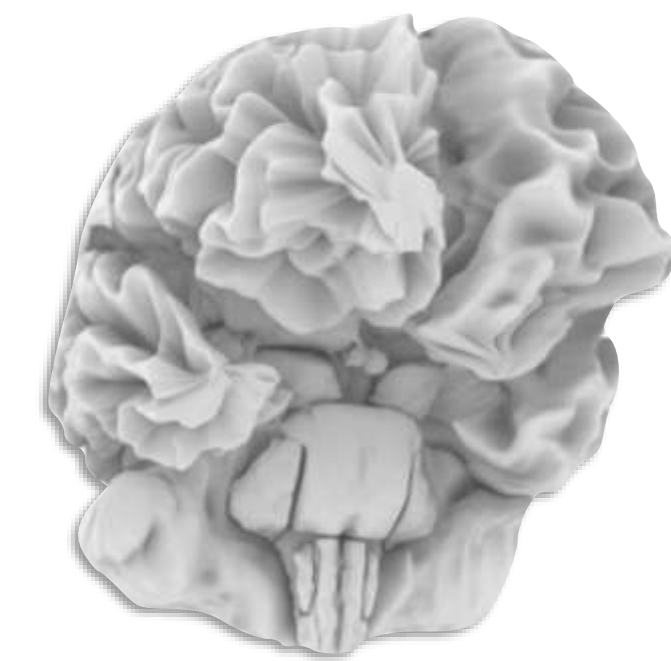


Full-ring CZT SPECT/CT

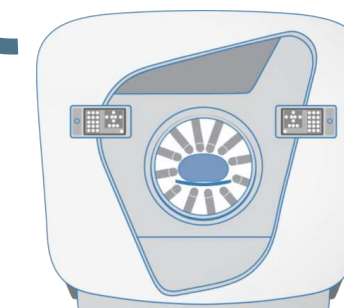
Allows the detectors to be positioned very close to the patient's head, providing **high sensitivity** and **improved spatial resolution**

Can **enhance image quality** as well as accuracy and precision of **quantitation** while reducing the acquisition time and/or injected dose [1-4]

Has the potential to enable **Cerebral Blood Flow (CBF)** quantification, thanks to **3D dynamic imaging capability**

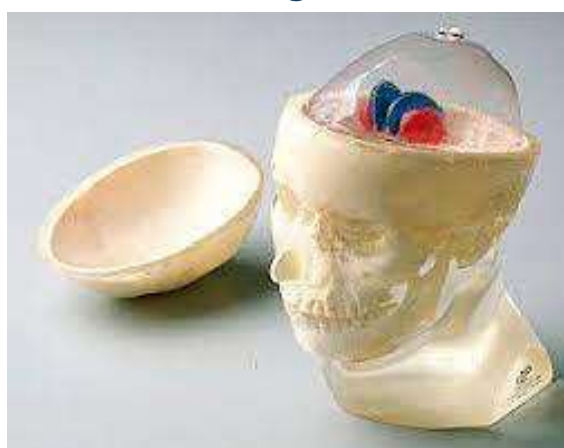


- [1] Bordonne, *et al.* *EJNMMI* 2020.
- [2] Piatkova, *et al.* *Clinic. Nucl. Med.* 2022.
- [3] Rogash, *et al.* *Nuklearmedizin-NuclearMedicine.* 2019.
- [4] Huh, *et al.* *Med. Phys.* 2021.



SPECT/CT Acquisition (¹²³I)

- Striatal Phantom study
- Dose: 3.8 mCi
- Str/Bkg ratio 8:1



- Comparison for an equal Scan Time (~3 min) against a conventional SPECT/CT with //hole LEHR collimator (Siemens Intevo)

Conventional	10 itr. 8 sub. Flash3D AC+SC
Full-Ring CZT	12 itr. 8 sub. PSFRq AC+SC

Coronal



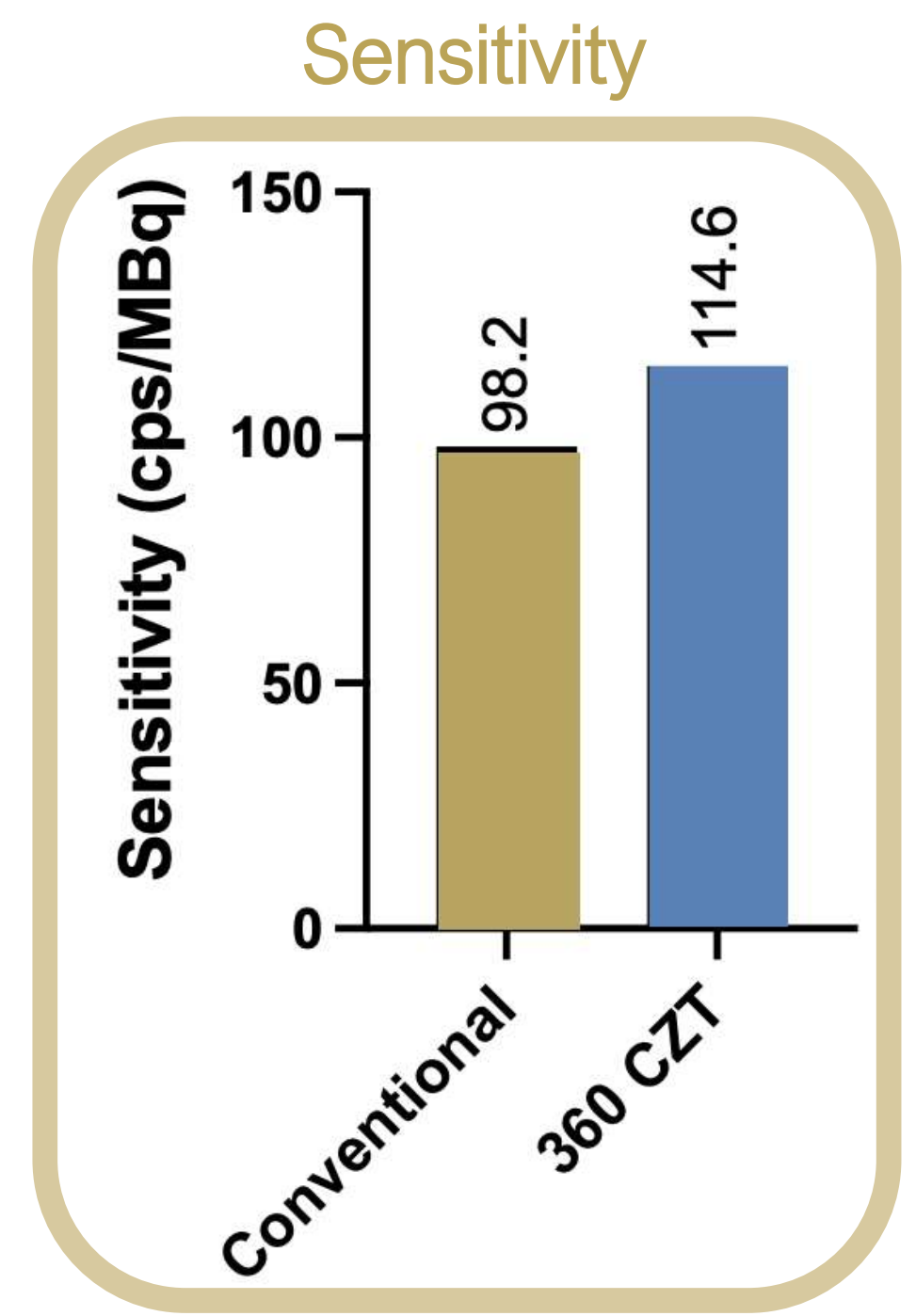
Sagittal



Transverse

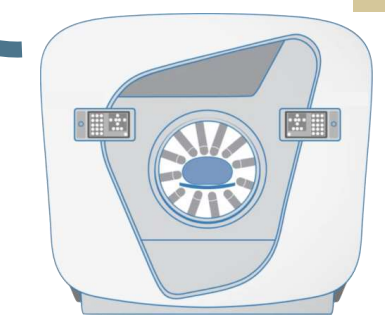


Posterior



Improved sensitivity (+16%), contrast, and overall IQ

Reduced penetration



SPECT/CT Acquisition (¹²³I)

- ❑ **Striatal Phantom** study
- ❑ Dose: 3.8 mCi
- ❑ Str/Bkg ratio 8:1

- ❑ Comparison for an equal **Scan Time (~3 min)** against a conventional SPECT/CT with //hole LEHR collimator (Siemens Intevo)

Conventional
 10 itr. 8 sub.
 Flash3D
 AC+SC

Coronal



Left

Sagittal



Posterior

Transverse

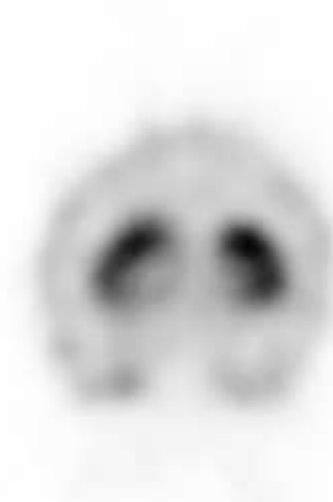


Left

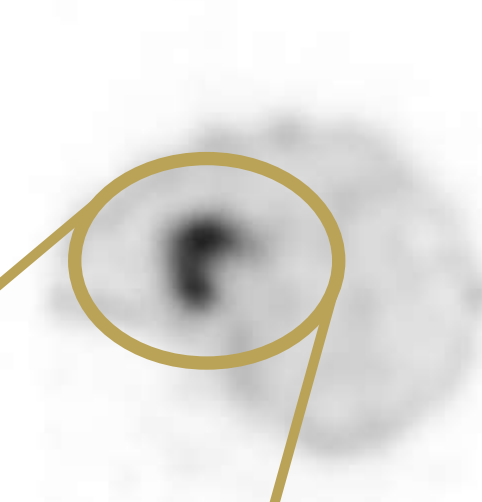
3D



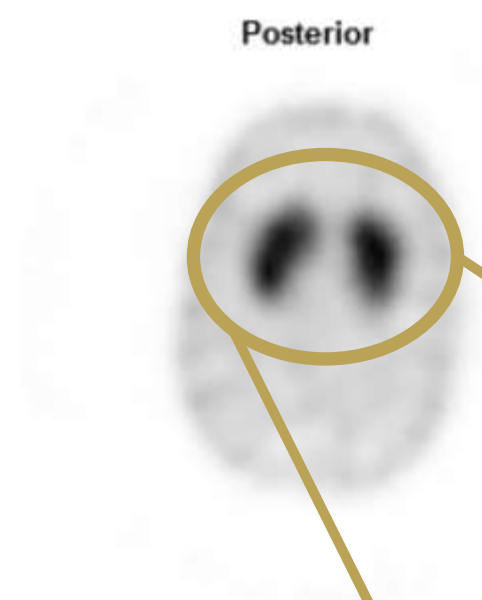
Full-Ring CZT
 12 itr. 8 sub.
 PSFRq
 AC+SC



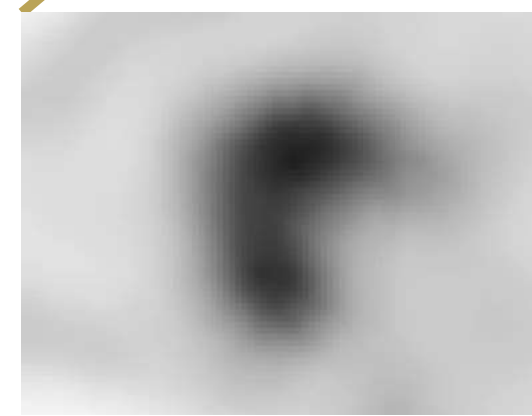
Left



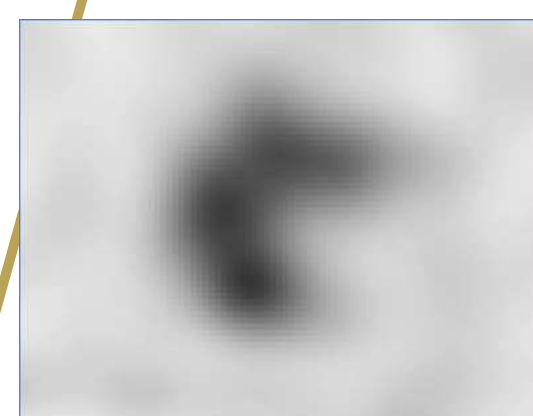
Posterior



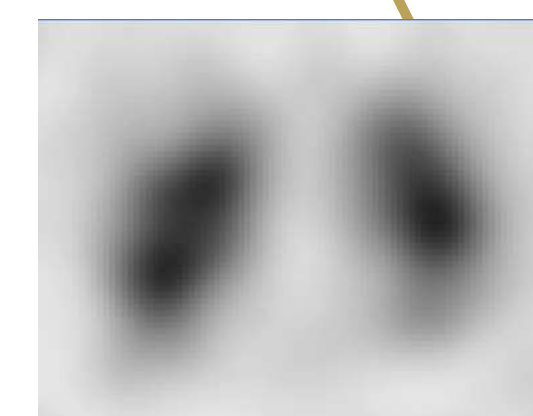
Left



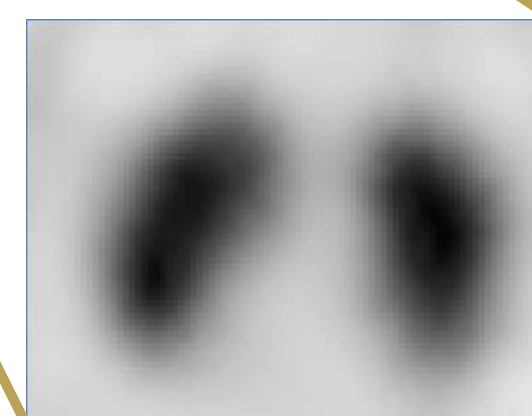
Full-Ring CZT



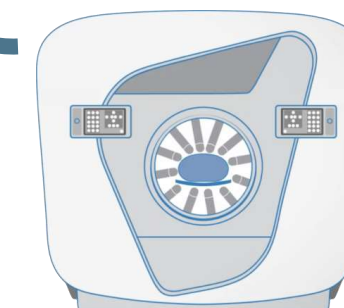
Conventional



Conventional



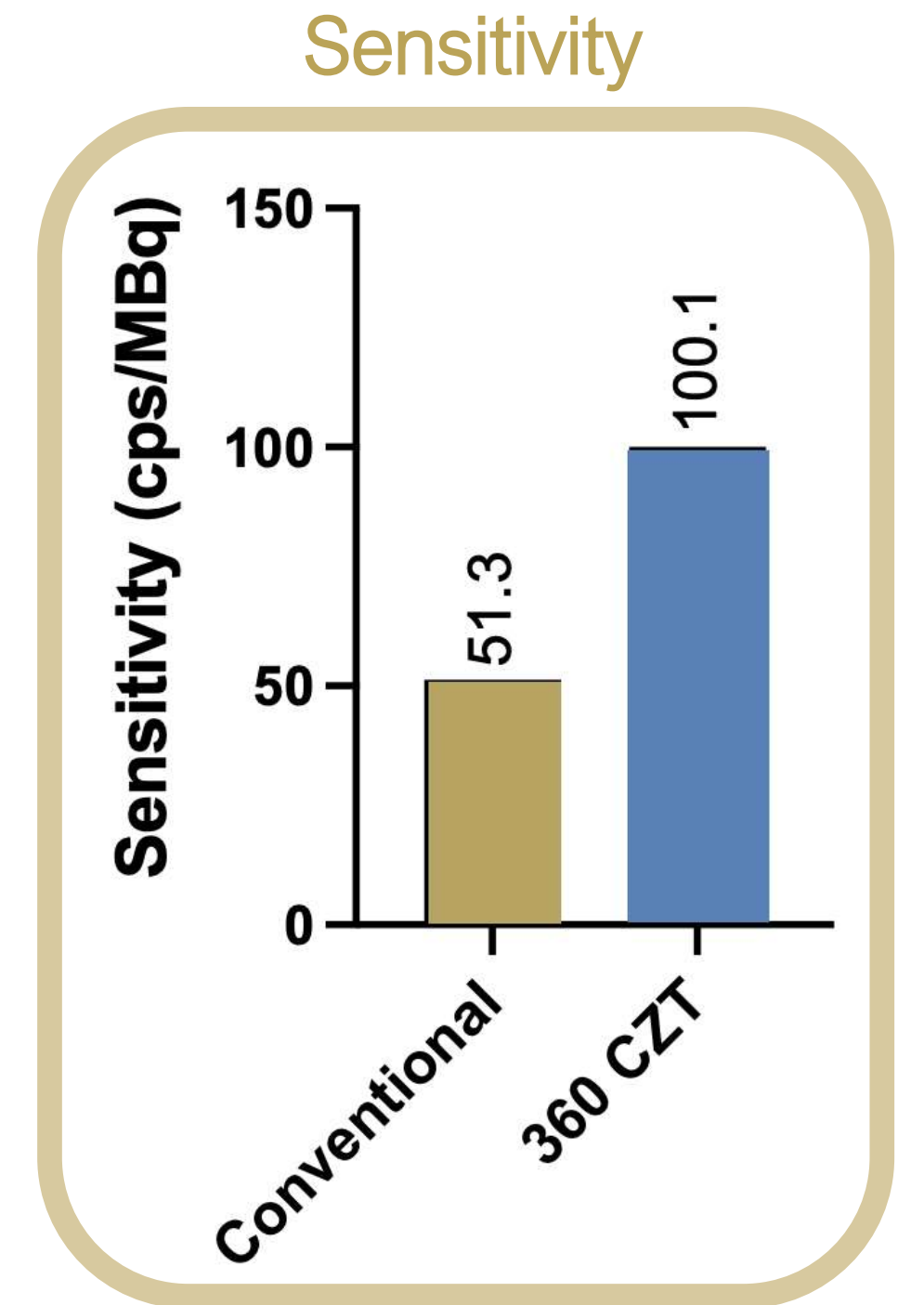
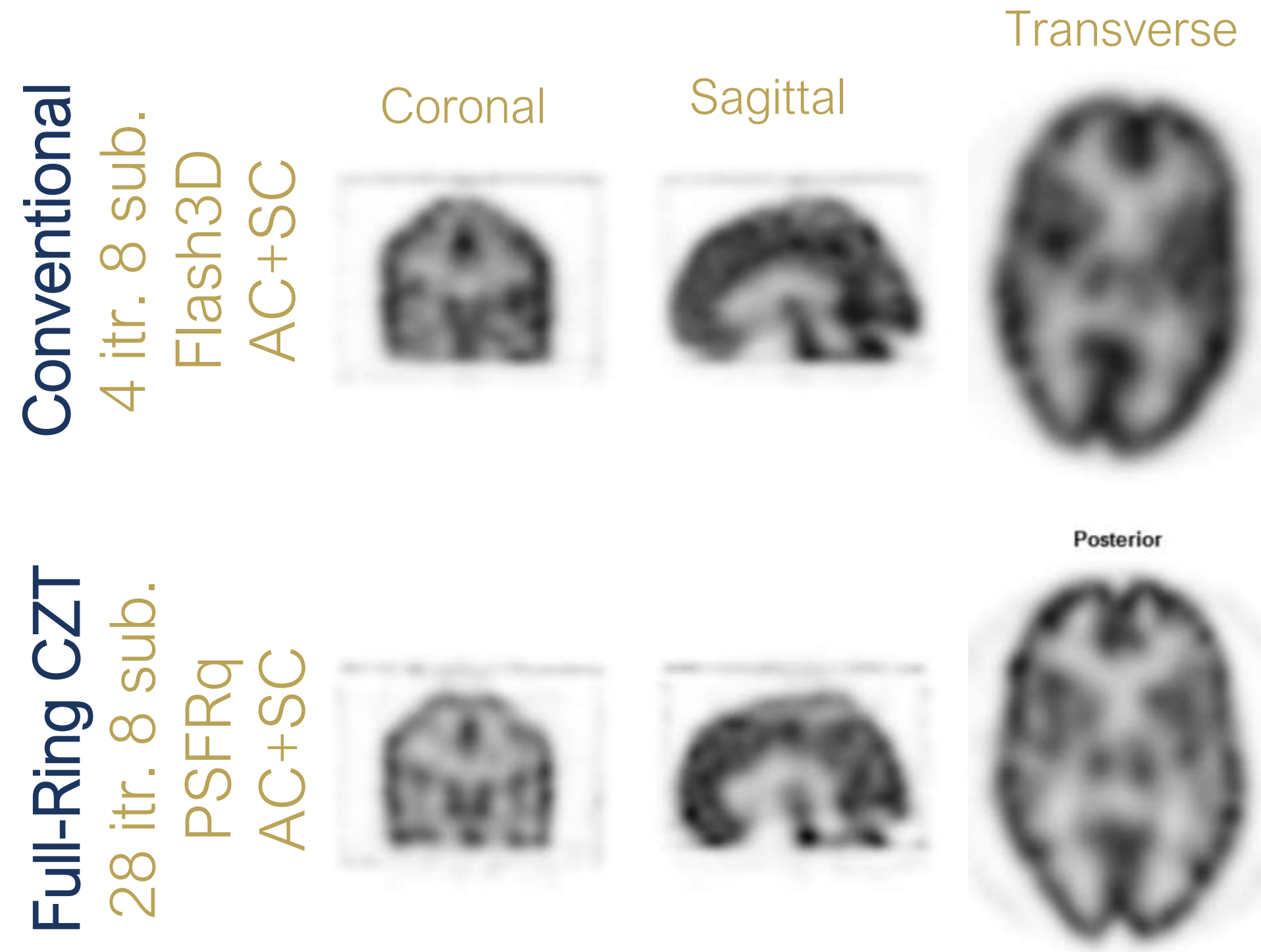
Full-Ring CZT



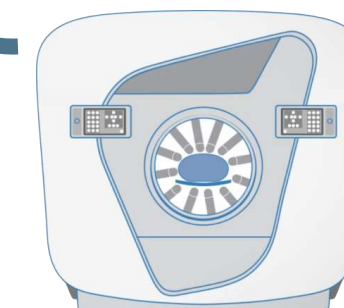
SPECT/CT Acquisition (^{99m}Tc)

- ❑ 3D Hoffman Phantom study
- ❑ Dose: 10.5 mCi
- ❑ Gray/White matter ratio 4:1

- ❑ Comparison for an equal Count Level (5.5 Mcts) against a conventional SPECT/CT with // -hole LEHR collimator (Siemens Intevo)

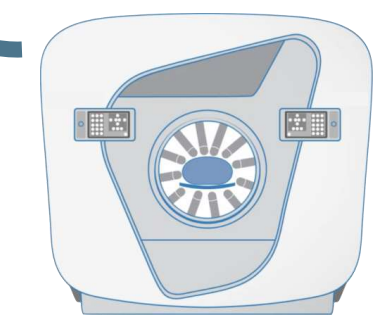
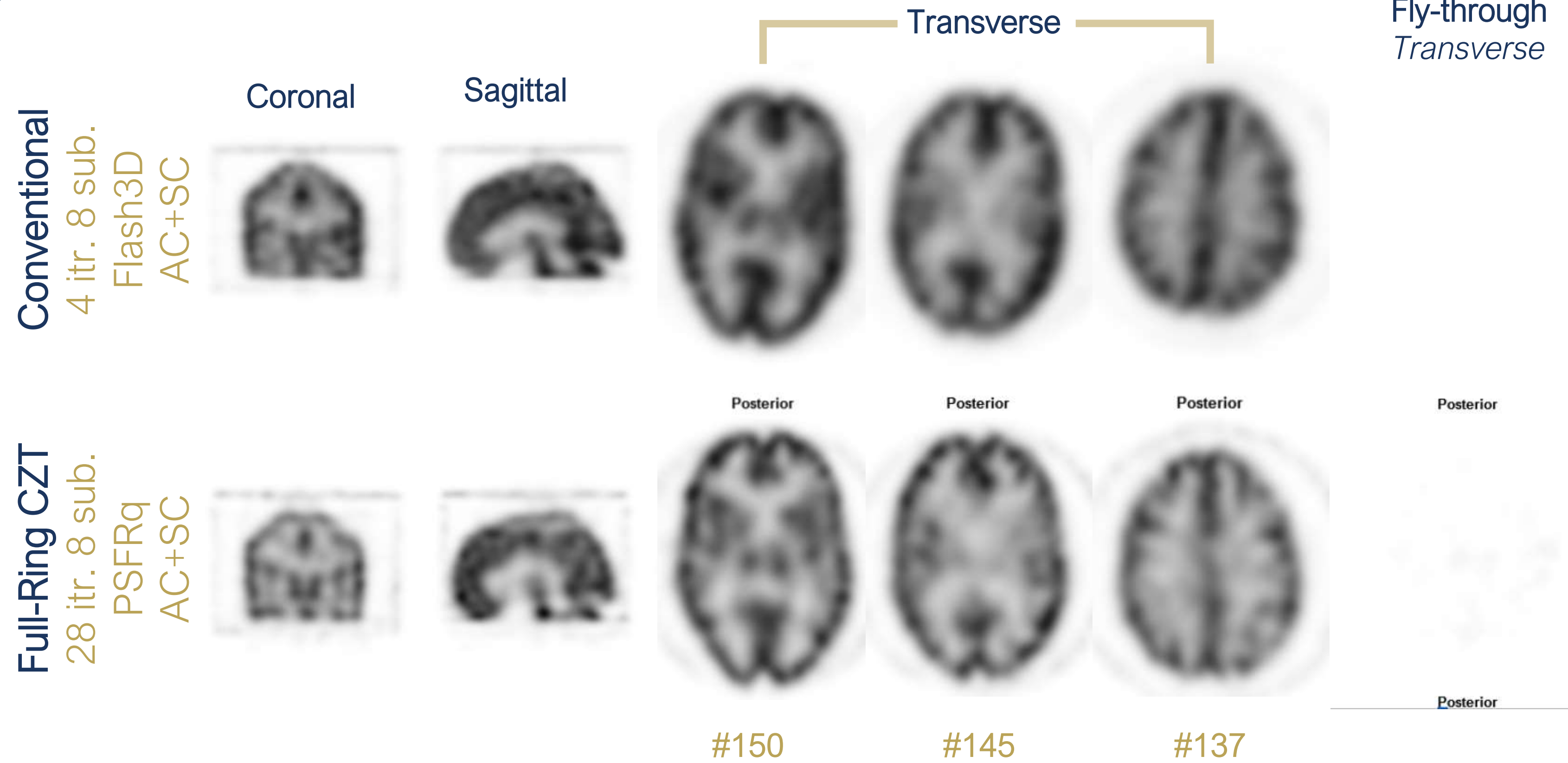


Improved sensitivity (2-fold), contrast, and overall IQ



SPECT/CT Acquisition (^{99m}Tc)

- 3D Hoffman Phantom study
- Dose: 10.5 mCi
- Gray/White matter ratio 4:1
- Comparison for an equal Count Level (5.5 Mcts) against a conventional SPECT/CT with //hole LEHR collimator (Siemens Intevo)

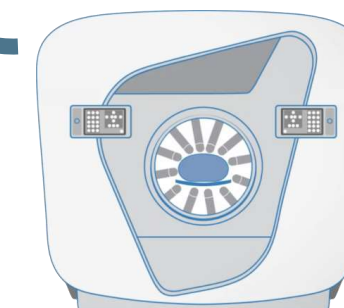
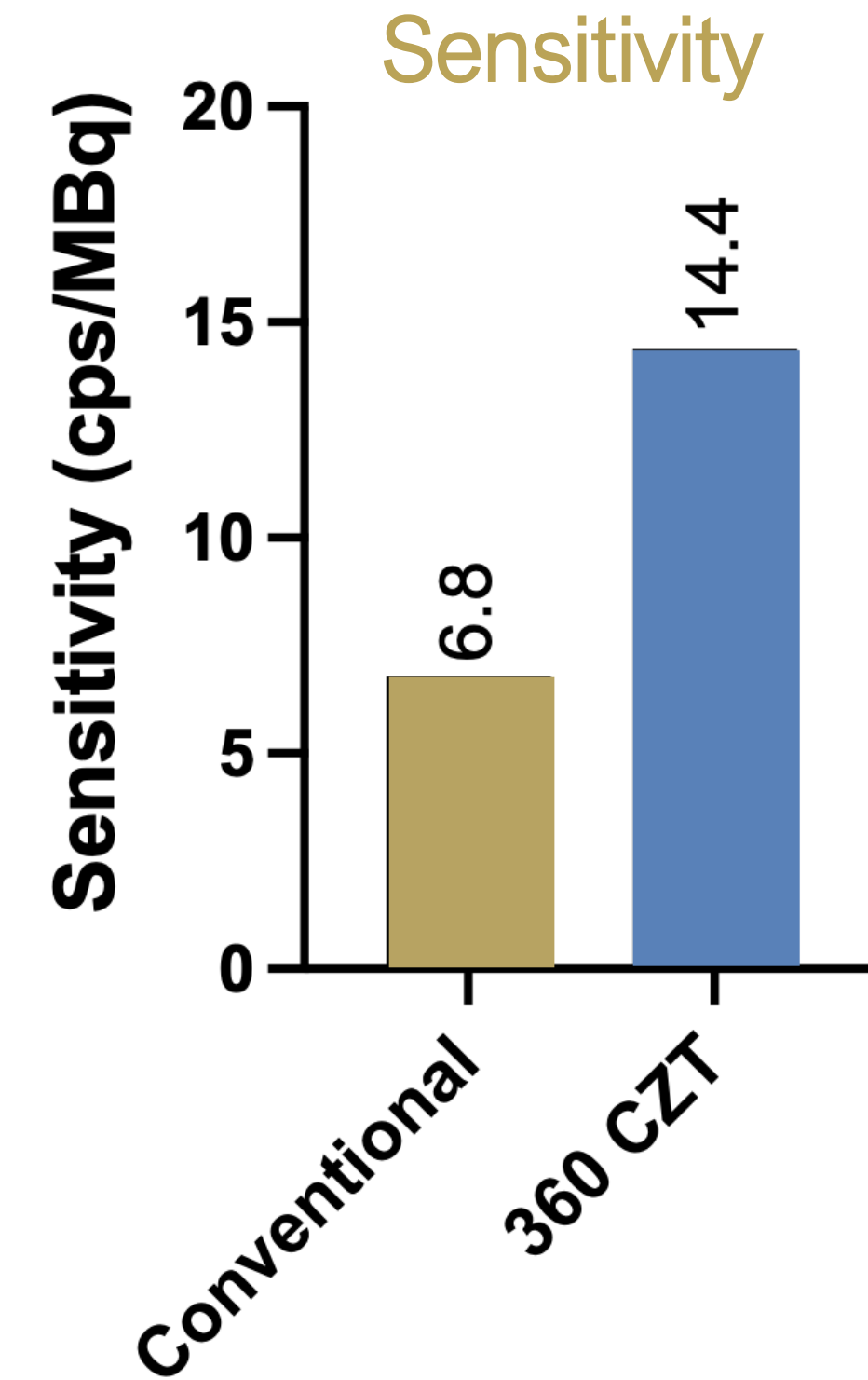
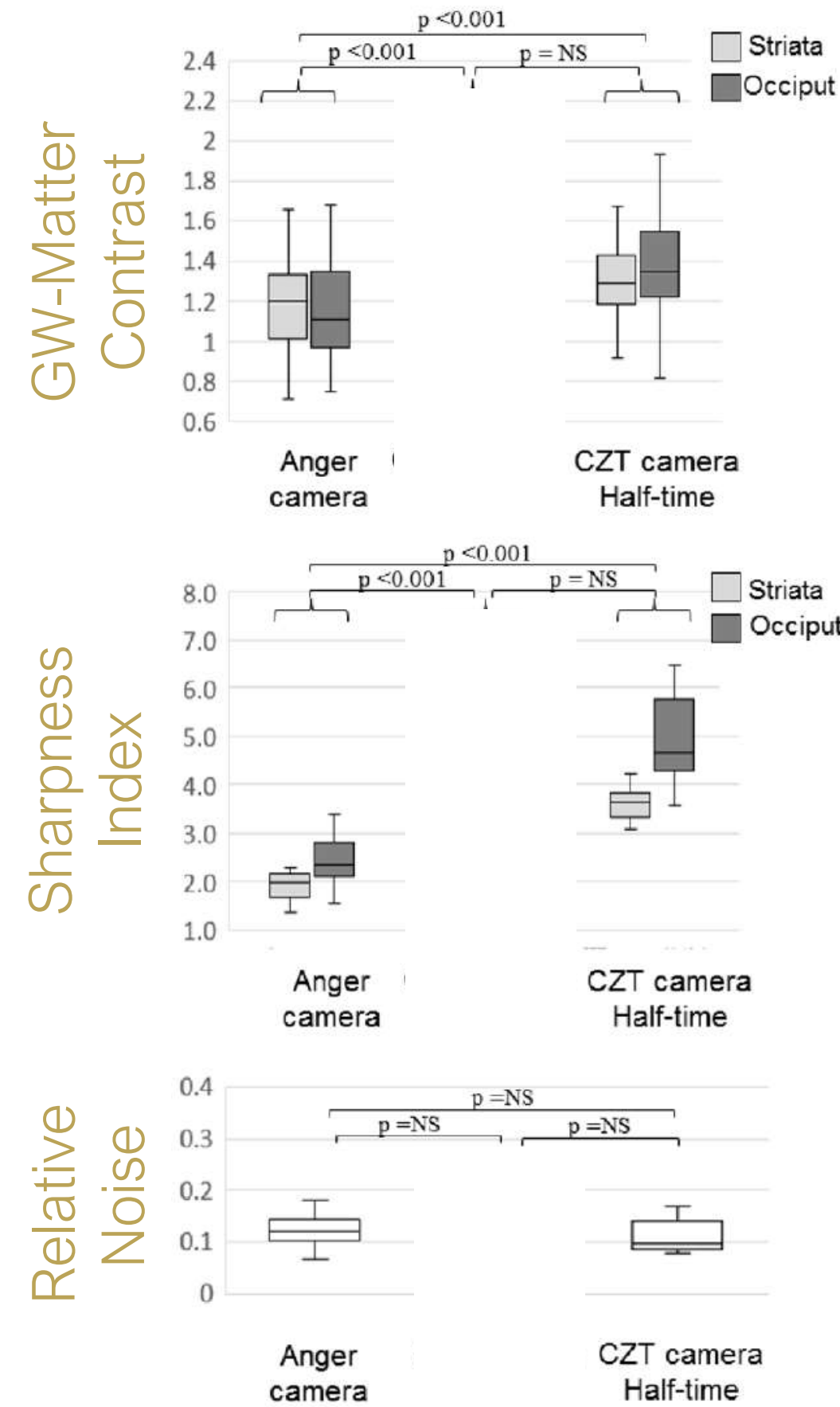
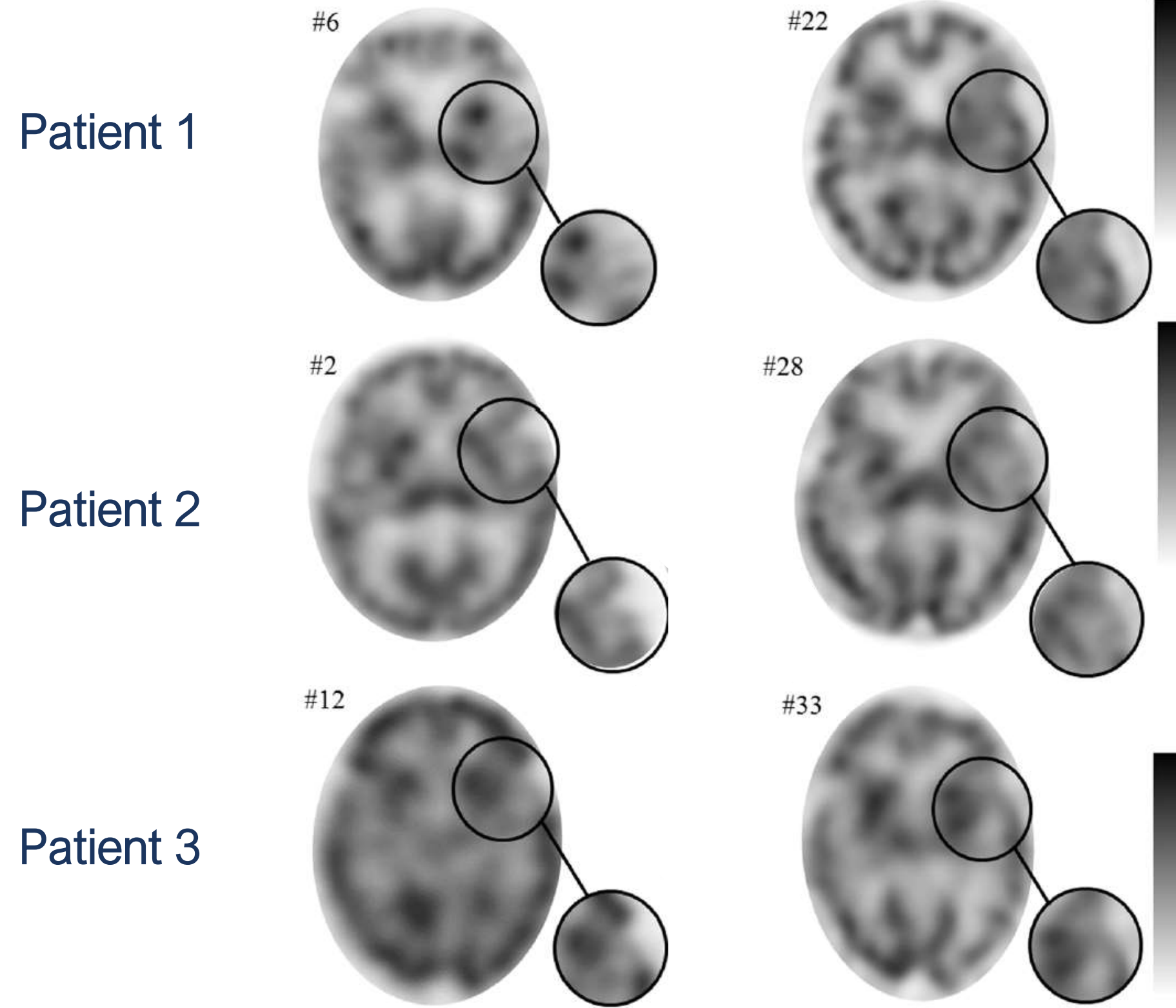


SPECT/CT Acquisition (^{99m}Tc-HMPAO) — Clinical Study

Adapted from Bordonne *et al.*
EJNMMI physics, 2020.

Conventional

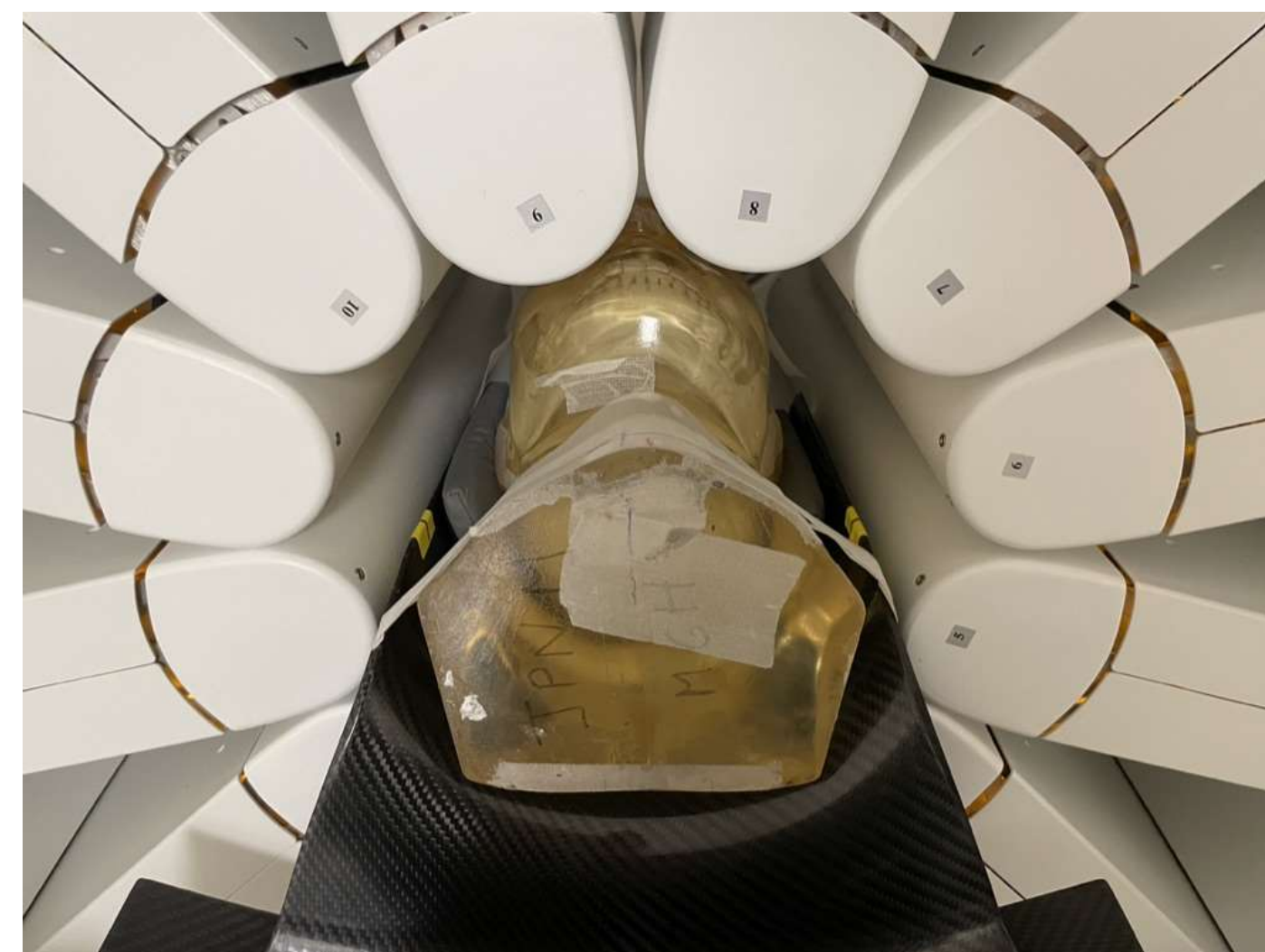
Full-Ring CZT



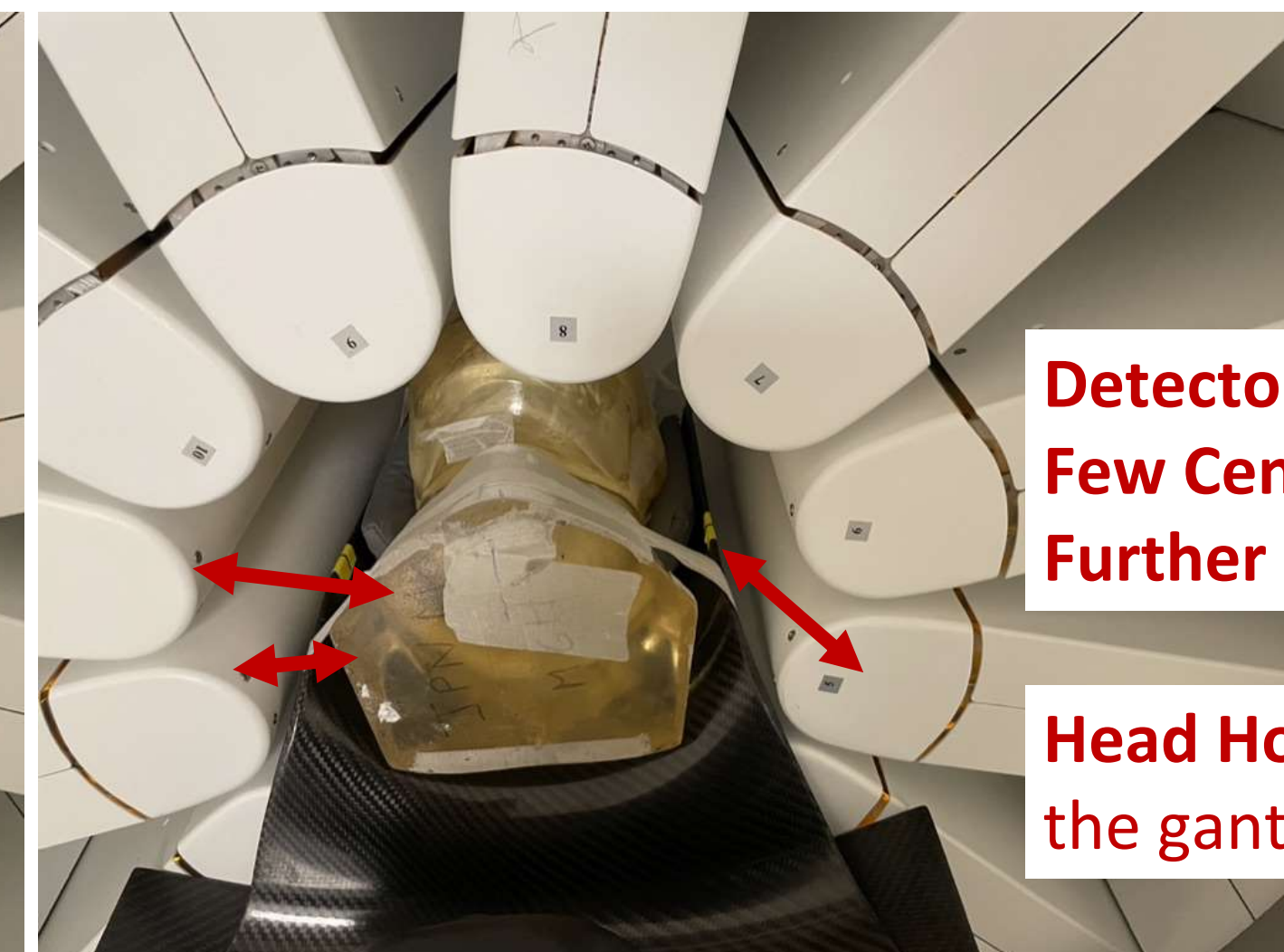
- Patient Positioning is of critical importance!**
- Detectors need to be positioned as close as possible to the brain for optimal imaging**
 Otherwise, **loss of spatial resolution!**
- Shoulders must be cleared.** Wider part of the **Head Holder** can also prevent detectors to be close enough!
- Need 1) for technologist training plus 2) to review data QC.**



Optimal Positioning



Sub-Optimal Positioning



Detectors are a Few Centimeters Further away!

Head Holder moved too far into the gantry!

Brain Perfusion SPECT/CT Acquisition (^{99m}Tc-ECD)

61y Male; Administrated Activity: 30 mCi

Sub-Optimal Positioning
Patient Too Far In the Gantry

Results in a Loss of Sp Resolution (*blurring*)

Optimal Positioning

Sharper!

Diamox 10/6

Baseline 10/18

The image displays two rows of SPECT/CT brain imaging results. The top row, labeled 'Sub-Optimal Positioning Patient Too Far In the Gantry', shows a blurred axial slice (frame 111) and a blurred sagittal slice (frame 150). The bottom row, labeled 'Optimal Positioning', shows a sharp axial slice (frame 20) and a sharp sagittal slice (frame 159). A red box highlights the sagittal slice in the optimal case, labeled 'Sharper!'. The text 'Results in a Loss of Sp Resolution (blurring)' is written in yellow above the top row, and 'Optimal Positioning' is written in white above the bottom row. The text 'Diamox 10/6' is written vertically on the left side of the top row, and 'Baseline 10/18' is written vertically on the left side of the bottom row. The text 'Sharper!' is written in red below the bottom row's sagittal slice.



Conventional SPECT suffers from,

- **Poor sensitivity** (*need to rotate to record data*),
- **Dependence on additional package or third-party software for absolute quantitation,**
- Being not really designed for **multi-bed SPECT Imaging**

Conventional SPECT Imaging relies on a mix of **planar 2D imaging** (*to identify the disease extent*) and **single or dual-bed SPECT** on areas of interest. Results in **Long Acquisition Time!**

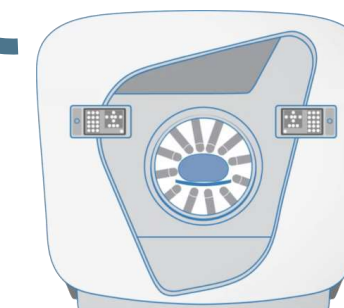
Full-Ring CZT SPECT alleviates these limitations by enabling **rapid whole-body SPECT/3D imaging**, plus offers **built-in quantitation [1-4]!**

[1] Melki, et al. EJNMMI 2020.

[2] Imbert, et al. J. Nucl. Med. 2019.

[3] Bahloul, et al. EJNMMI, 2023

[4] Song, et al. EJNMMI. 2023.



Full-Ring CZT SPECT/CT

Provides **high sensitivity** and **improved spatial resolution** that enhance **image quality** and **quantitation** while reducing the acquisition time and/or injected dose compared to conventional SPECT/CT [1-4]

Enables **rapid 3D whole-body imaging**

Improves **image contrast** for bone lesions compared to inherently limited planar imaging, which results in better localization of abnormal accumulations [1-4]



Whole Body SPECT/CT

- ☐ Scan time ~ **20 min**
- ☐ 59y Female
- ☐ BMI: 23.8
- ☐ ^{99m}Tc-HMDP 11.7 mCi

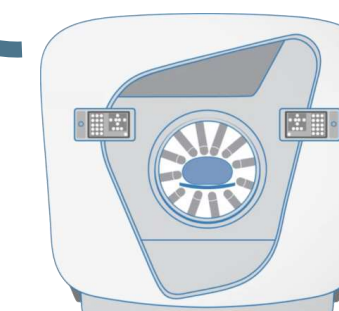
Images courtesy of Dept of Clinical
Physiology and Nuclear Medicine at
Linköping University Hospital,
Region Östergötland, Sweden

[1] Goshen, *et al. EJNMMI Phys.* 2018.

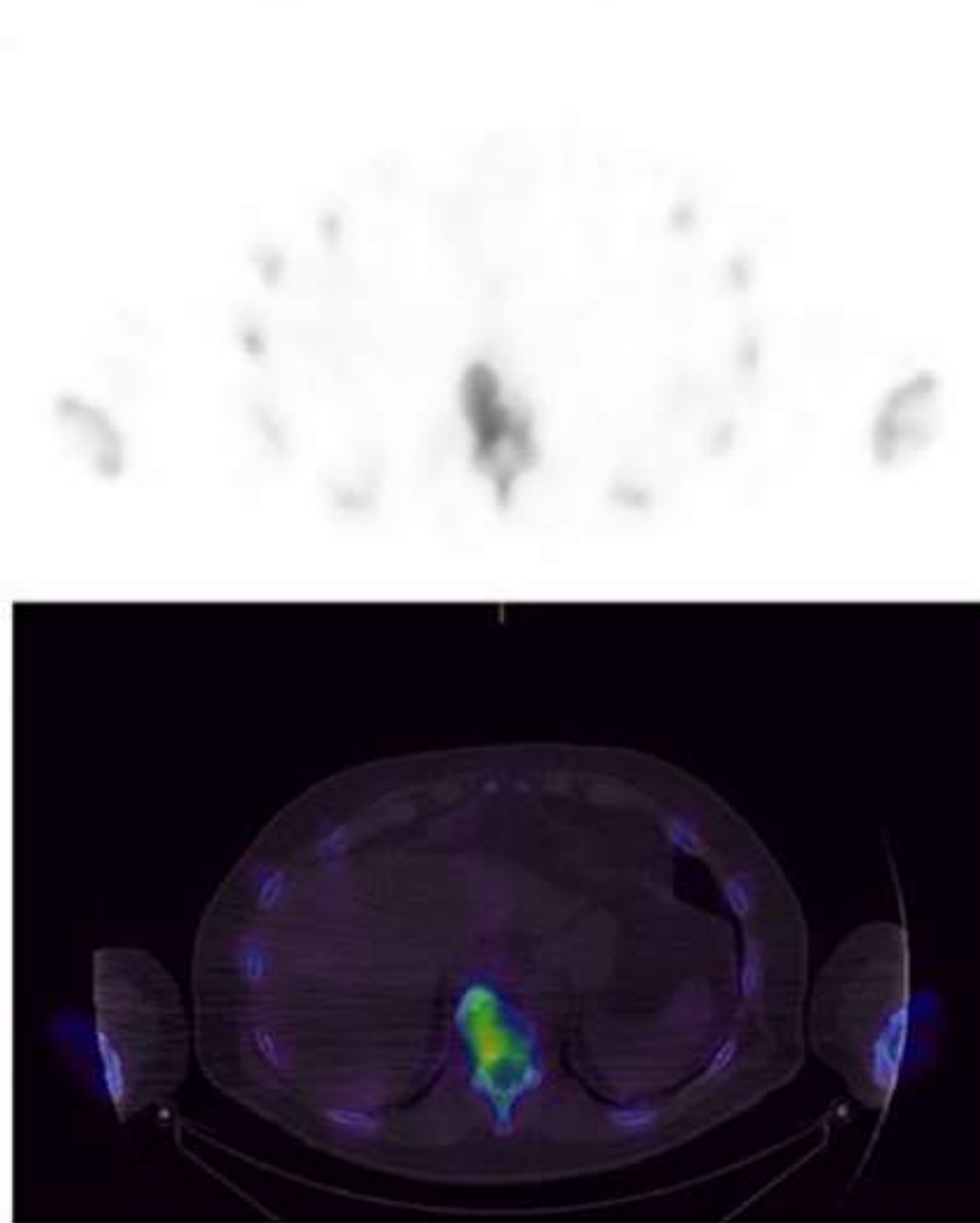
[2] Melki, *et al. Eur. J. Nucl. Med. Mol. Imaging* 2020.

[3] Imbert, *et al. J. Nucl. Med.* 2019.

[4] Yamane, *et al. Eur. J. Nucl. Med. Mol. Imaging.* 2019.

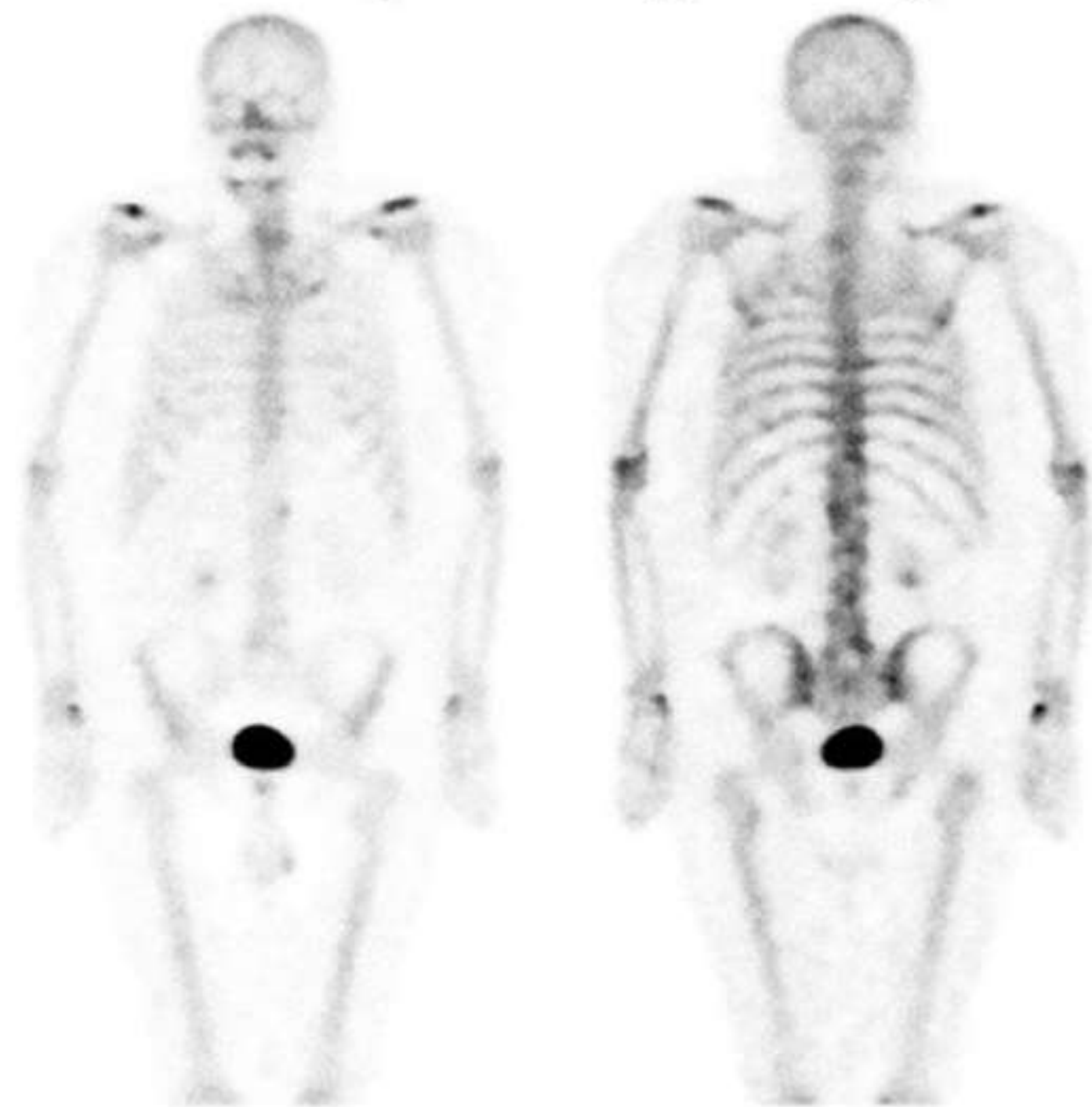


Starguide SPECT/CT



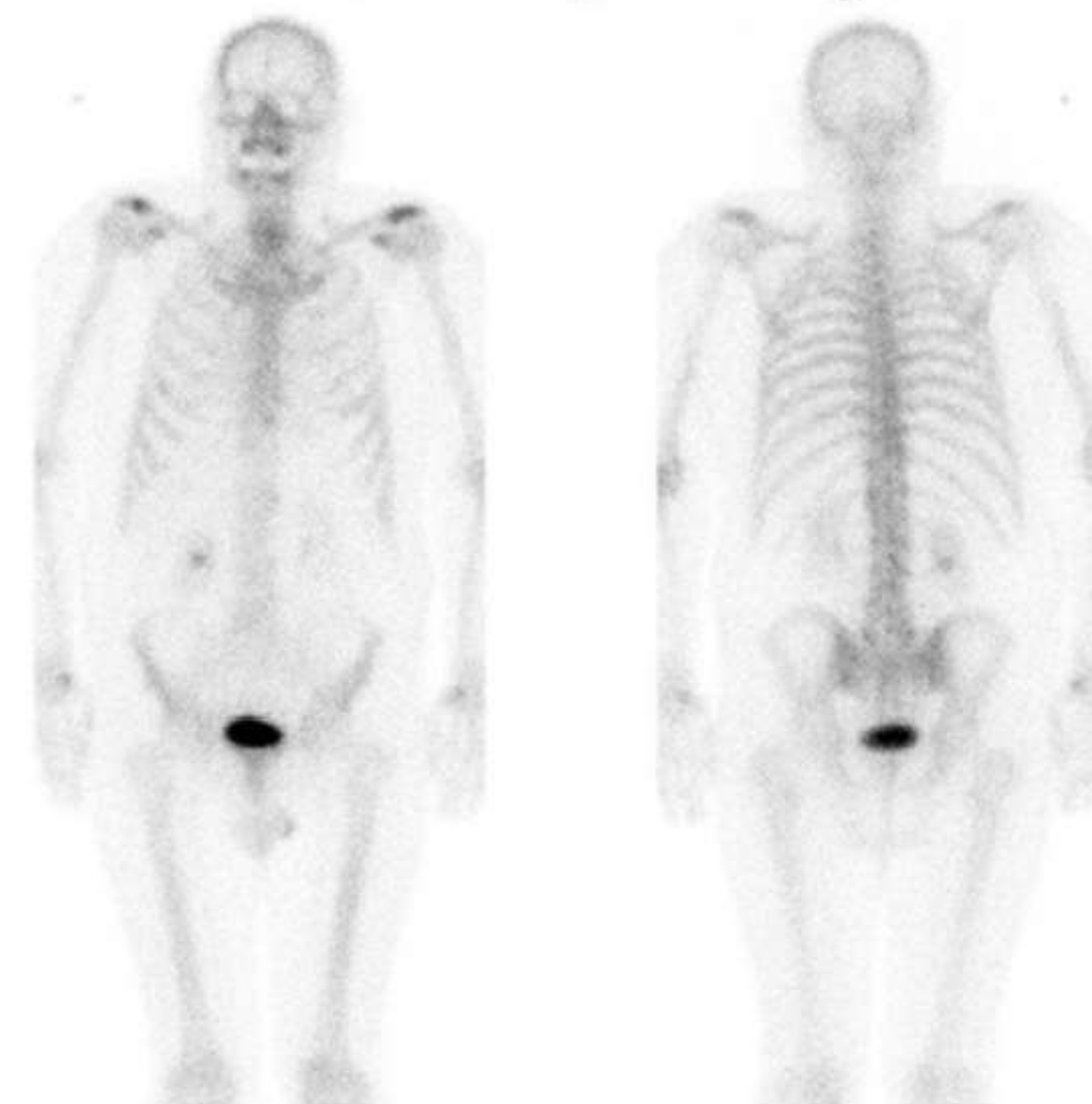
WB 3D SPECT

Pseudo-planar Starguide images



Synthetic 2D planar created from WB SPECT image

Conventional planar images



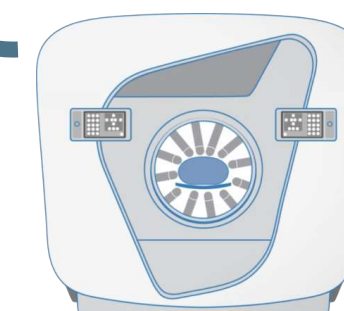
2D Planar Images

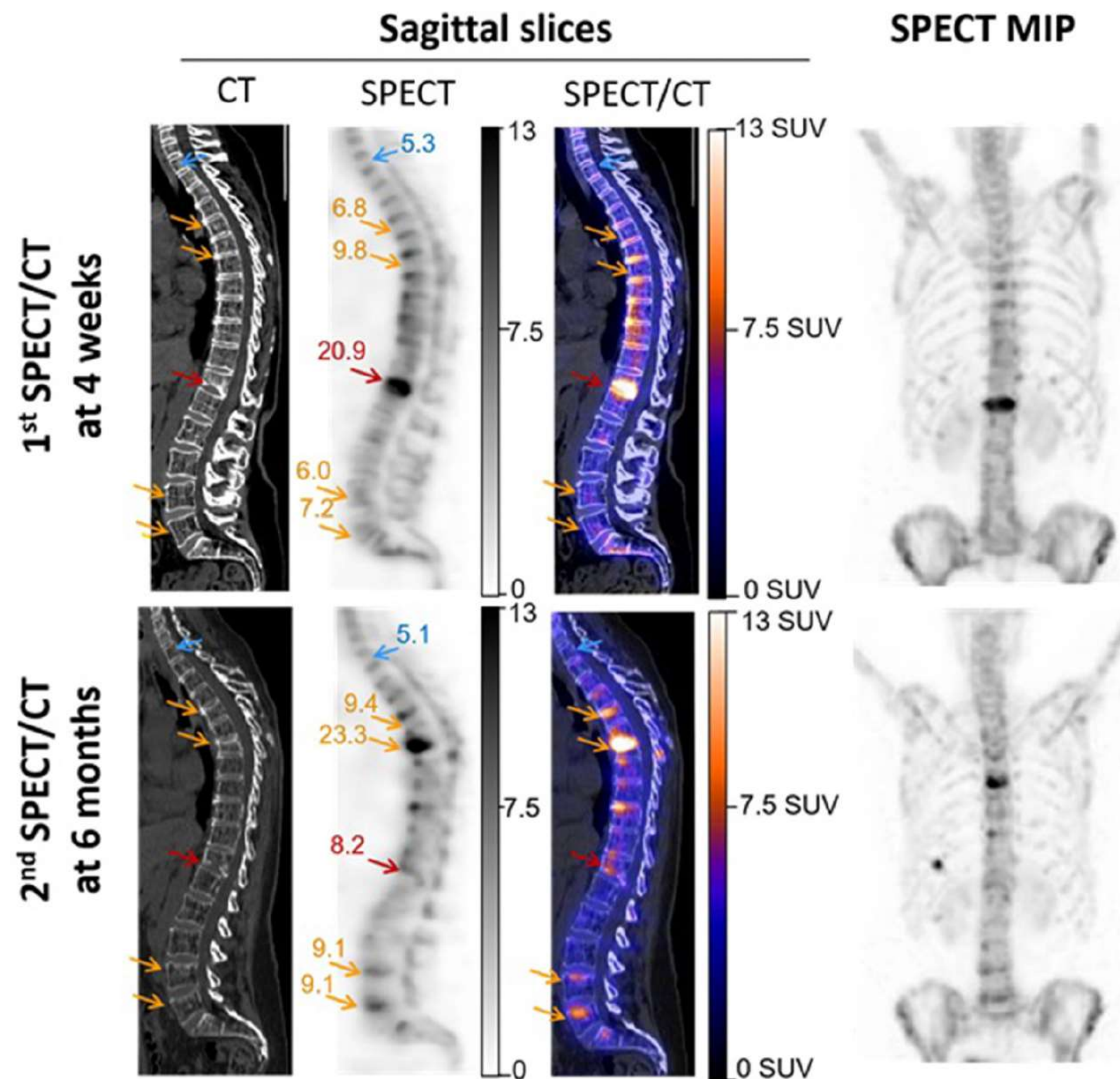
Scan Time ~ **20 min**

Scan Time = **15 min**
 Speed = 16 cm/min

Cerić Andelius, I., Minarik, D., Persson, E., Mosén, H., Valind, K., Trägårdh, E., & Oddstig, J. (2024). First clinical experience of a ring-configured cadmium zinc telluride camera: A comparative study versus conventional gamma camera systems. *Clinical Physiology and Functional Imaging*, 44(1), 79-88.

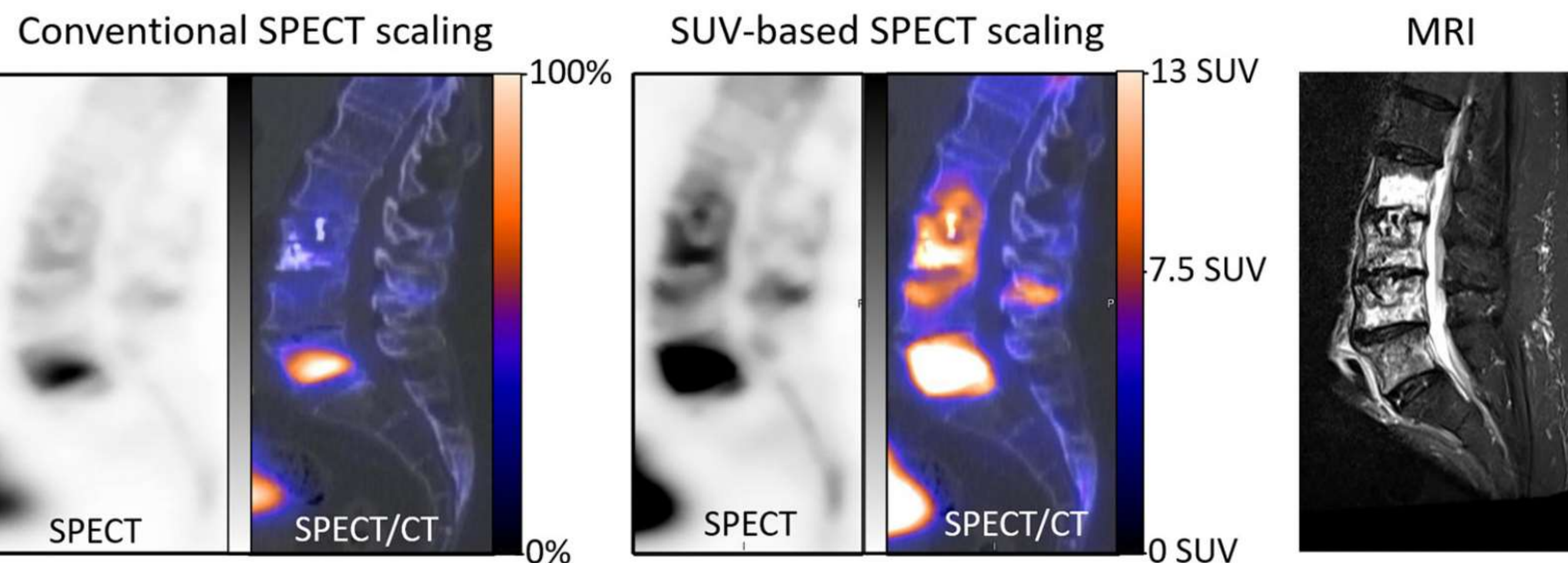
Usually followed by **Single-bed SPECT**
 (additional **20-30 min**)



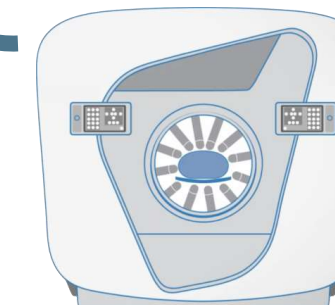


Full-Ring CZT SPECT/CT

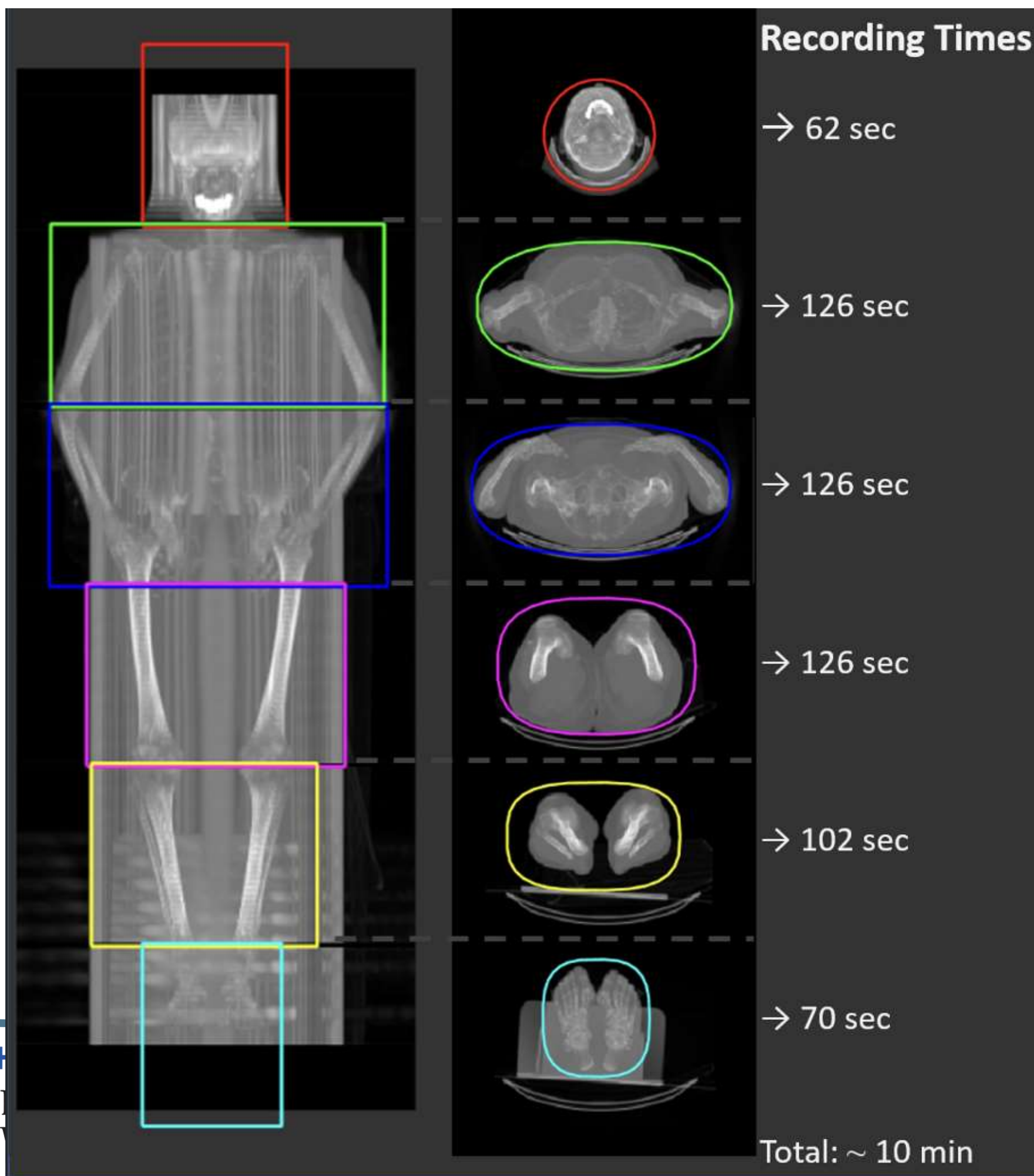
- Rapid Imaging 12-15 min scan time to acquire the entire spin
- Enabled-SUV quantitation to localize fracture



Adapted from Bahloul, A., Verger, A., Blum, A., Chawki, M. B., Perrin, M., Melki, S., et al. (2021). Bone scintigraphy of vertebral fractures with a whole-body CZT camera in a PET-like utilization. *Frontiers in Nuclear Medicine*, 1, 740275.



Body Focus Mode

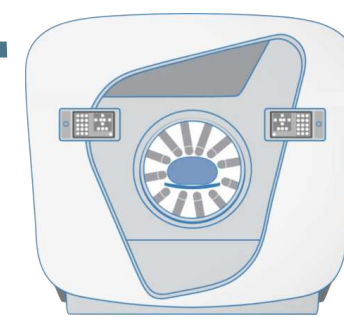


- Modulation of the **detector contouring per bed**. Focus VOIs derived from CT

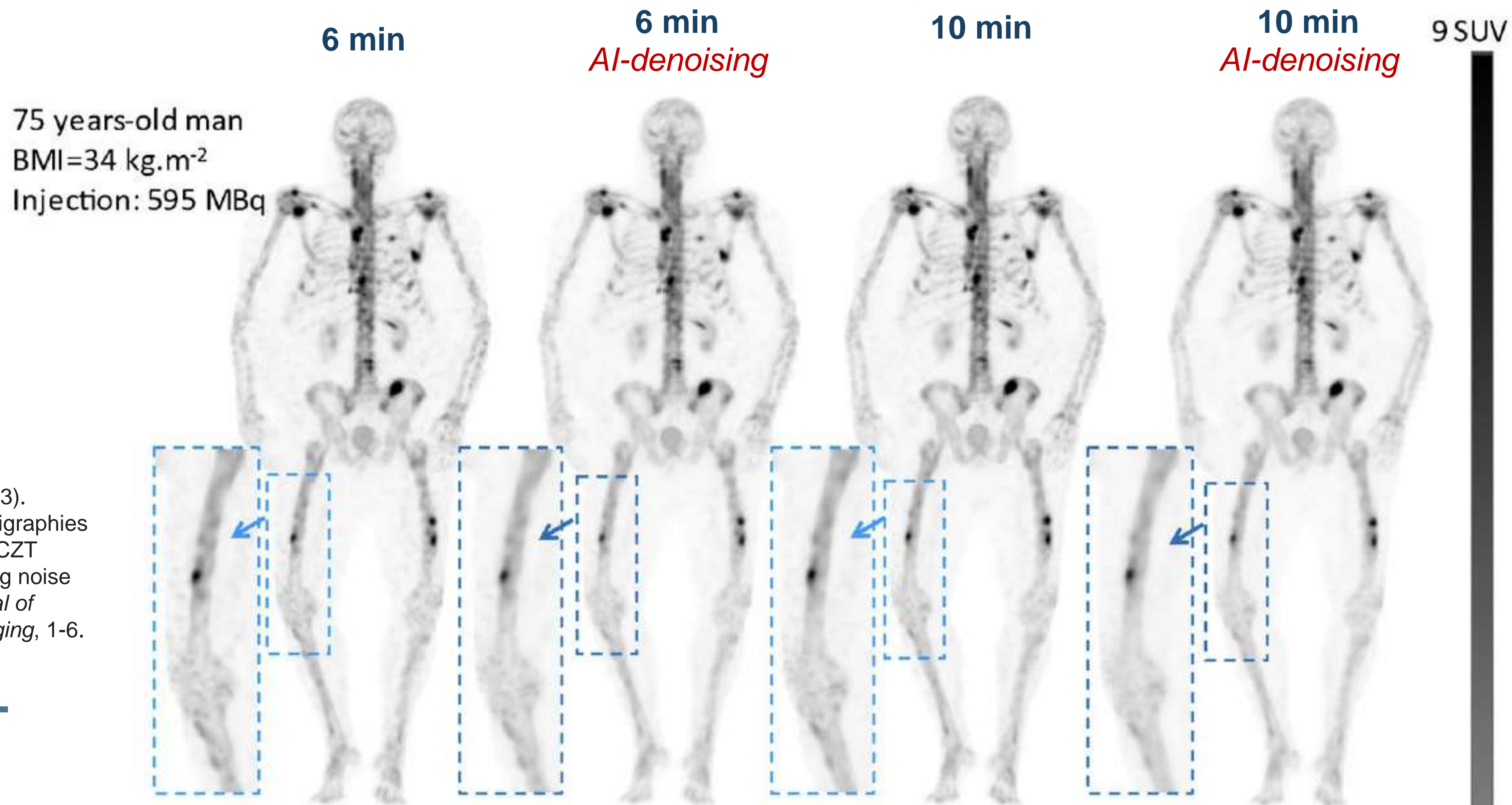
- Provides a **uniform scan time per millimeter**. Noise equivalent image per bed.

- Improvement in sensitivity resulting in **further scan time reduction (15-20 min down to 10 min)**

Adapted from Bahloul, A., Verger, A., Lamash, Y., Roth, N., Dari, D., Marie, P. Y., & Imbert, L. (2023). Ultra-fast whole-body bone tomoscintigraphies achieved with a high-sensitivity 360° CZT camera and a dedicated deep-learning noise reduction algorithm. *European Journal of Nuclear Medicine and Molecular Imaging*, 1-6.



Further scan time reduction can be achieved with **AI-denoising** from **10 min** down to **6 min** (*1 min/bed*)



Adapted from Bahloul, A., et al. (2023). Ultra-fast whole-body bone tomoscintigraphies achieved with a high-sensitivity 360° CZT camera and a dedicated deep-learning noise reduction algorithm. *European Journal of Nuclear Medicine and Molecular Imaging*, 1-6.

Full-ring CZT SPECT/CT

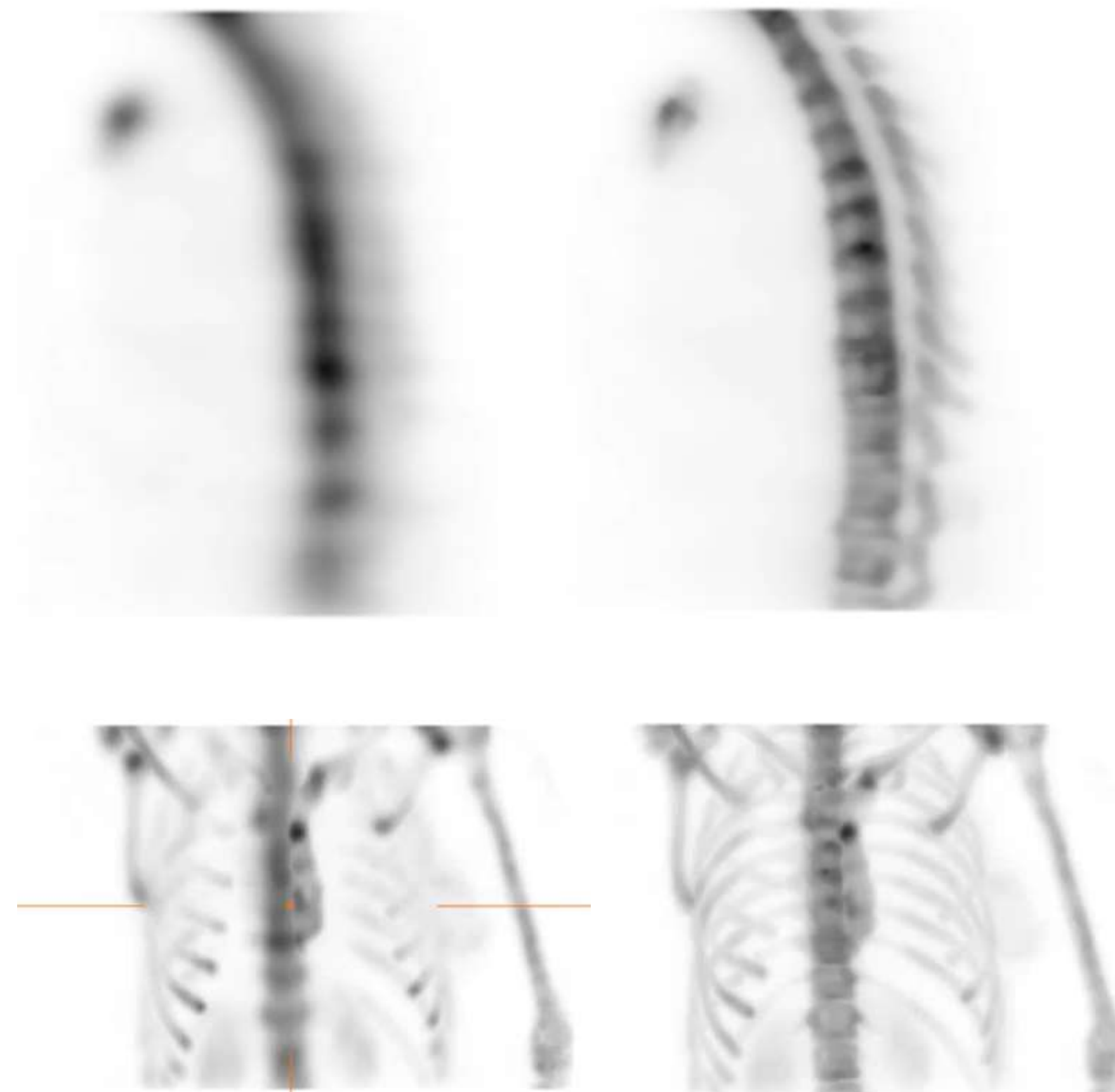
Provides high sensitivity and improved spatial resolution that enhance image quality and quantitation while reducing the acquisition time and/or injected dose [1-4]

Improves image contrast for bone lesions compared to inherently limited planar imaging, which results in better localization of abnormal accumulations [1-4]

Can provide **CT-based anatomic information** that can be used for **Partial Volume Correction (PVC)** to further improve **image quality** and **quantitation** compared to SPECT-only and planar imaging

w/o PVC

w/ PVC



[1] Goshen, *et al. EJNMMI Phys.* 2018.

[2] Melki, *et al. Eur. J. Nucl. Med. Mol. Imaging* 2020.

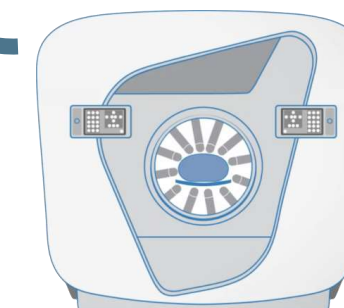
[3] Imbert, *et al. J. Nucl. Med.* 2019.

[4] Yamane, *et al. Eur. J. Nucl. Med. Mol. Imaging.* 2019.



- Rapid growth of targeted radionuclide therapies**, such as ^{177}Lu -Dotatate and ^{177}Lu -PSMA [1,2]
- Major Interest in personalized dosimetry and treatment response assessment and monitoring** [3,4]
- SPECT** has major advantages compared to PET for image-based dosimetry due its **capability to image directly** ^{177}Lu
- Growing need for high-speed whole-body SPECT/CT**. Conventional SPECT is limited by long acquisition because of poor sensitivity [5]
- Full-Ring SPECT/CT** can overcome these challenges thanks to its **improved imaging performance**

[1] Sartor, *et al.* *N Engl J Med.* 2021.
[2] Strosberg, *et al.* *N Engl J Med.* 2017.
[3] Sgouros, *et al.* *J. Nucl. Med.* 2021.
[4] Pandit-Taskar, *et al.* *J. Nucl. Med.* 2021.
[5] Pathmanandavel, *et al.* *J. Nucl. Med.* 2022.



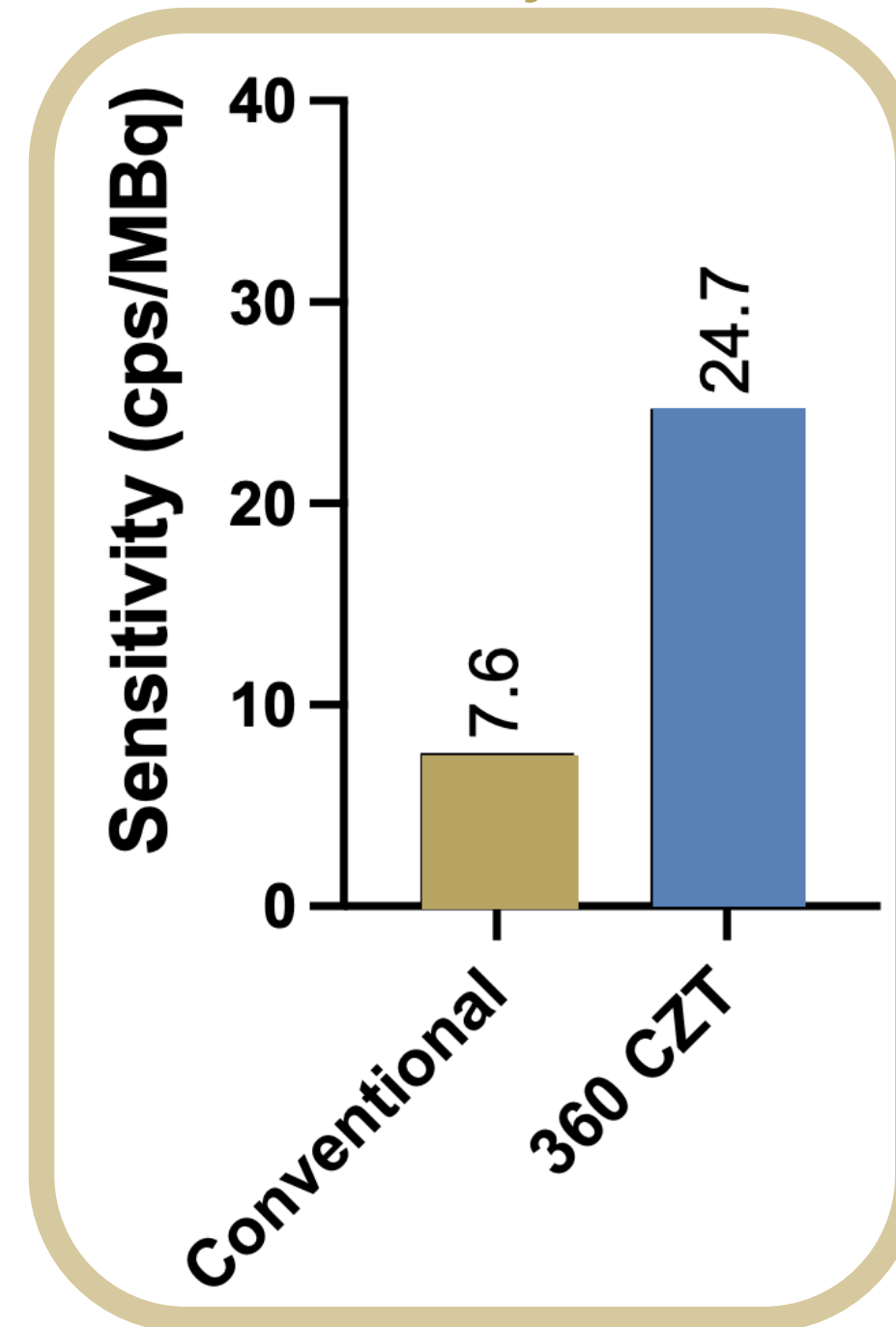
SPECT/CT Acquisition (¹⁷⁷Lu)

- ☐ Uniform Phantom study
- ☐ Dose: 30 mCi
- ☐ Volume: 6415 ml

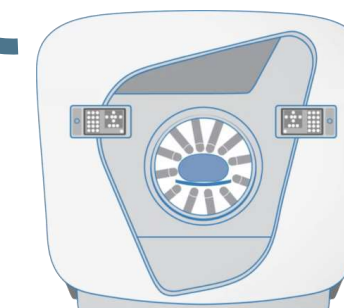
- ☐ **Conventional**
 Siemens Intevo w/ MEGP
 208 keV ± 10 %
 Upper/Lower Scatter windows 10%
 CTAC

- ☐ **Full-Ring CZT**
 Veriton 200 Series
 113 keV ± 10 %
 Upper/Lower Scatter windows 10%
 CTAC

Sensitivity for a uniform cylinder



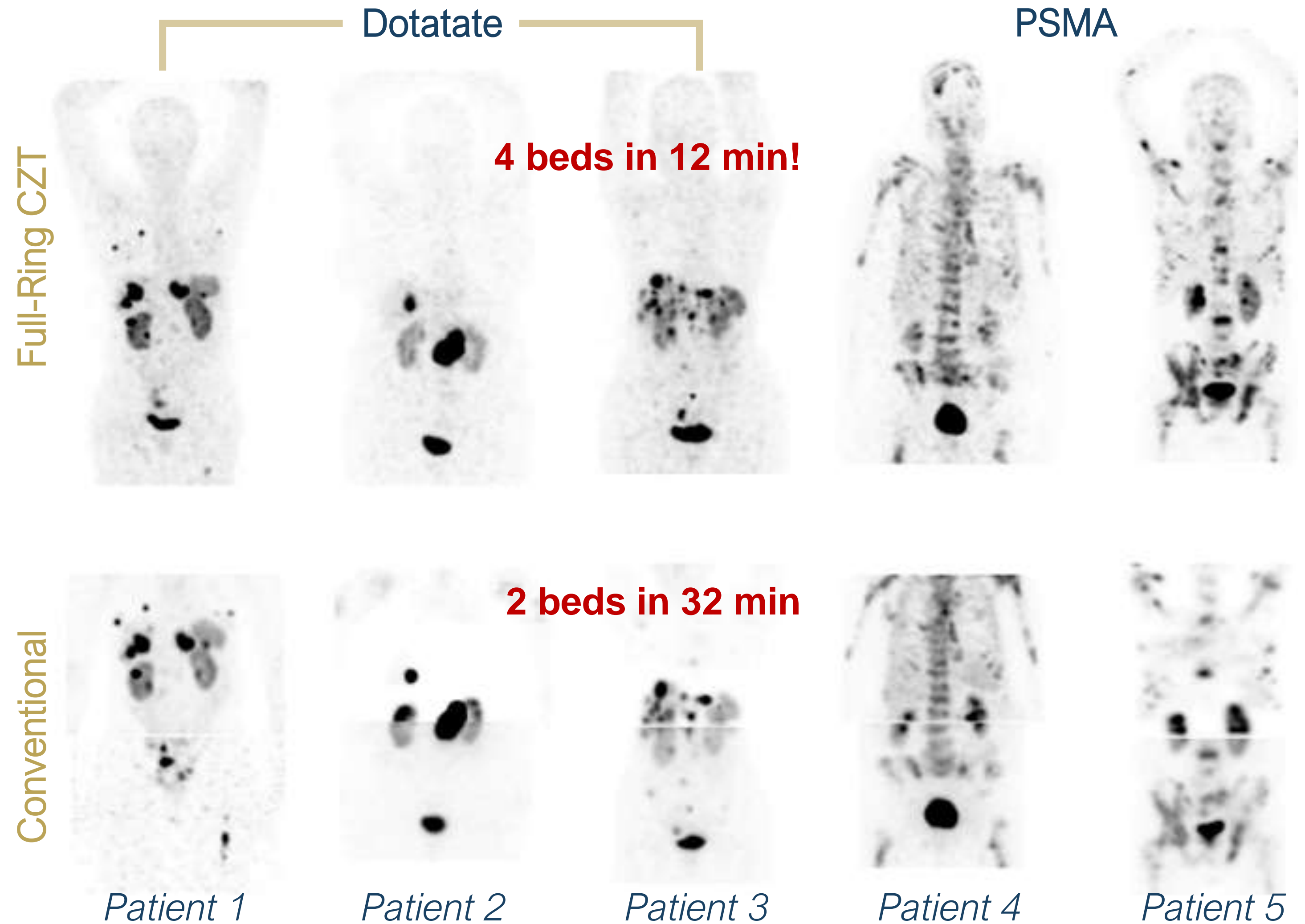
- Improvement in sensitivity of 3-fold for the least abundant ¹⁷⁷Lu-peak
- Further enhancement with Veriton 400 series and Starguide capable to image both peaks of ¹⁷⁷Lu



SPECT/CT Acquisition

- **Full-Ring CZT**
 GE StarGuide
 113 keV ± 10 % & 208 keV ± 6 %
 Upper/Lower Scatter windows 10%
 CTAC
 4 bed positions / **12 min** total scan

- **Conventional**
 GE 670 Pro w/ MEGP
 113 keV ± 10 % & 208 keV ± 6 %
 Upper/Lower Scatter windows 10%
 CTAC
 2 bed positions / **32 min** total scan



Total Body SPECT/CT Acquisition (¹⁷⁷Lu)

- ❑ 67y Male
- ❑ BMI: 22.6

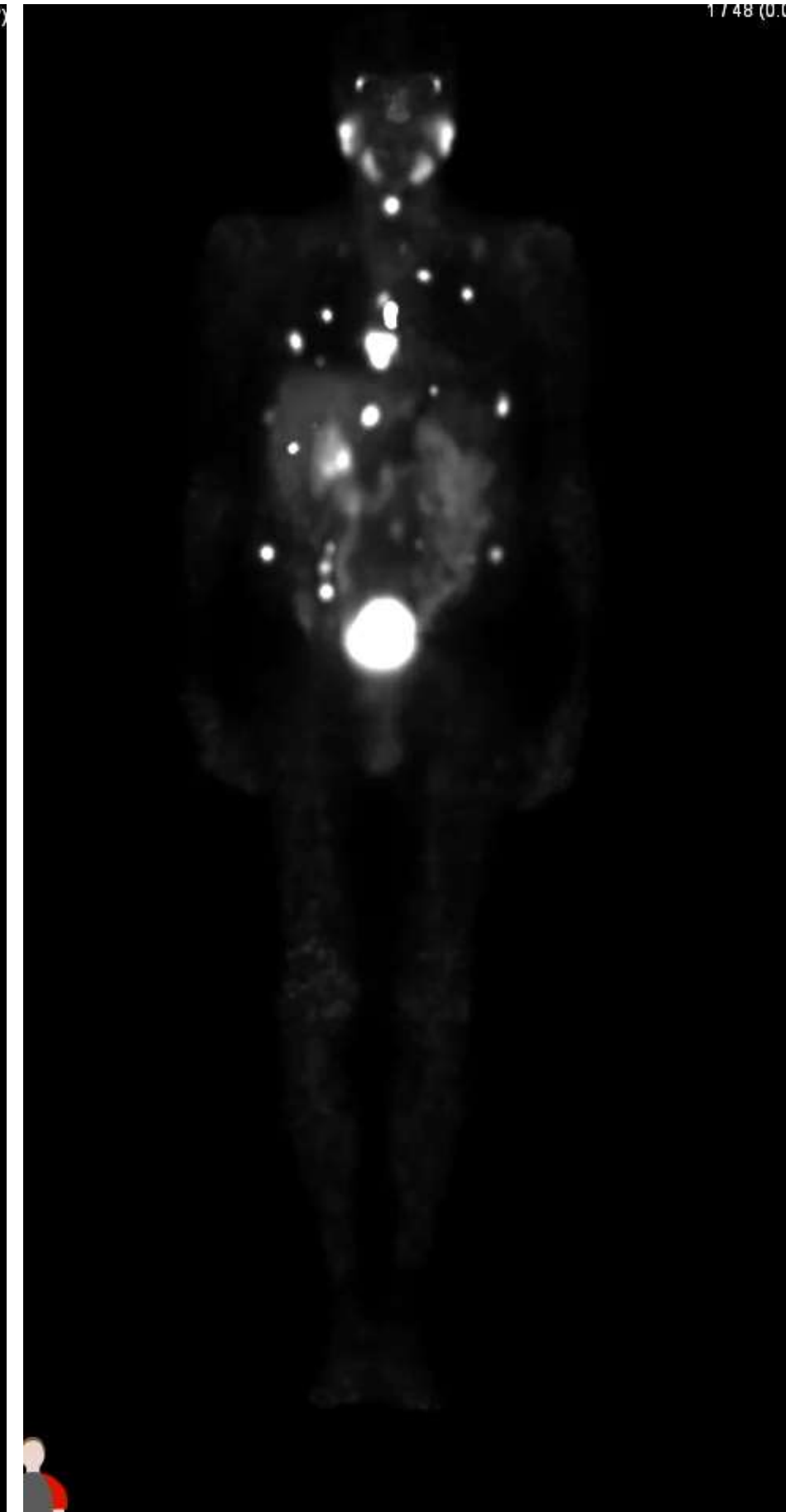
- ❑ Dose: 202.3 mCi of ¹⁷⁷Lu-PSMA
- ❑ Acquisition time: **18 min**

- ❑ **Full-Ring CZT**
 Veriton 200 Series
 113 keV ± 10 %
 Upper/Lower Scatter windows 10%
 CTAC

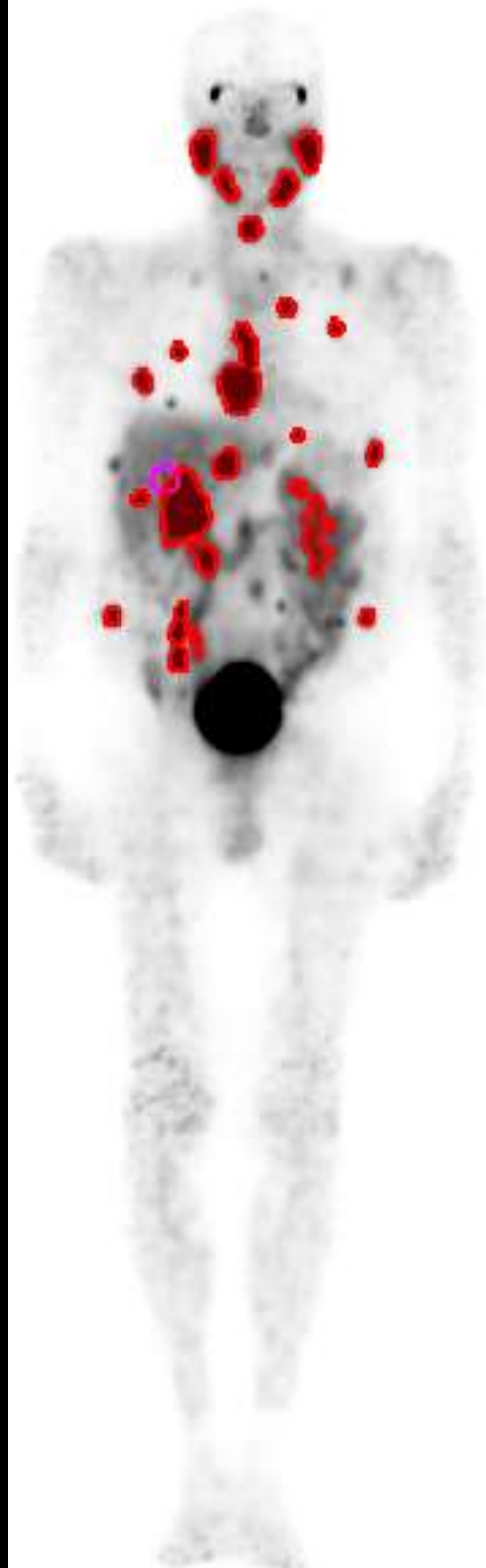
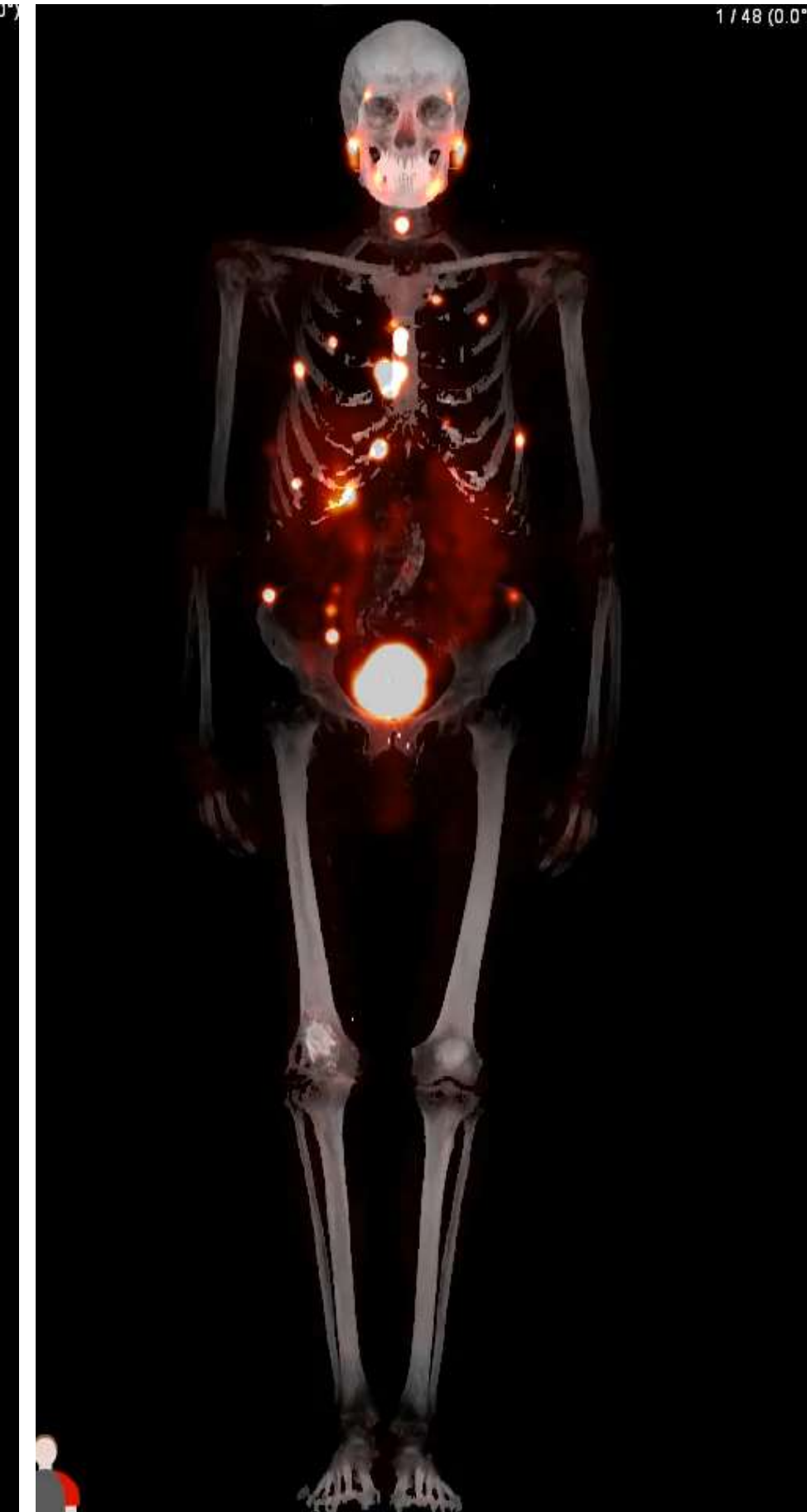
CT



SPECT

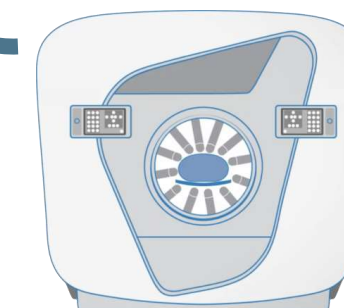


SPECT/CT



18 min

Images courtesy of Centre Hospitalier Régional Universitaire (CHRU) de Nancy, Nancy, France

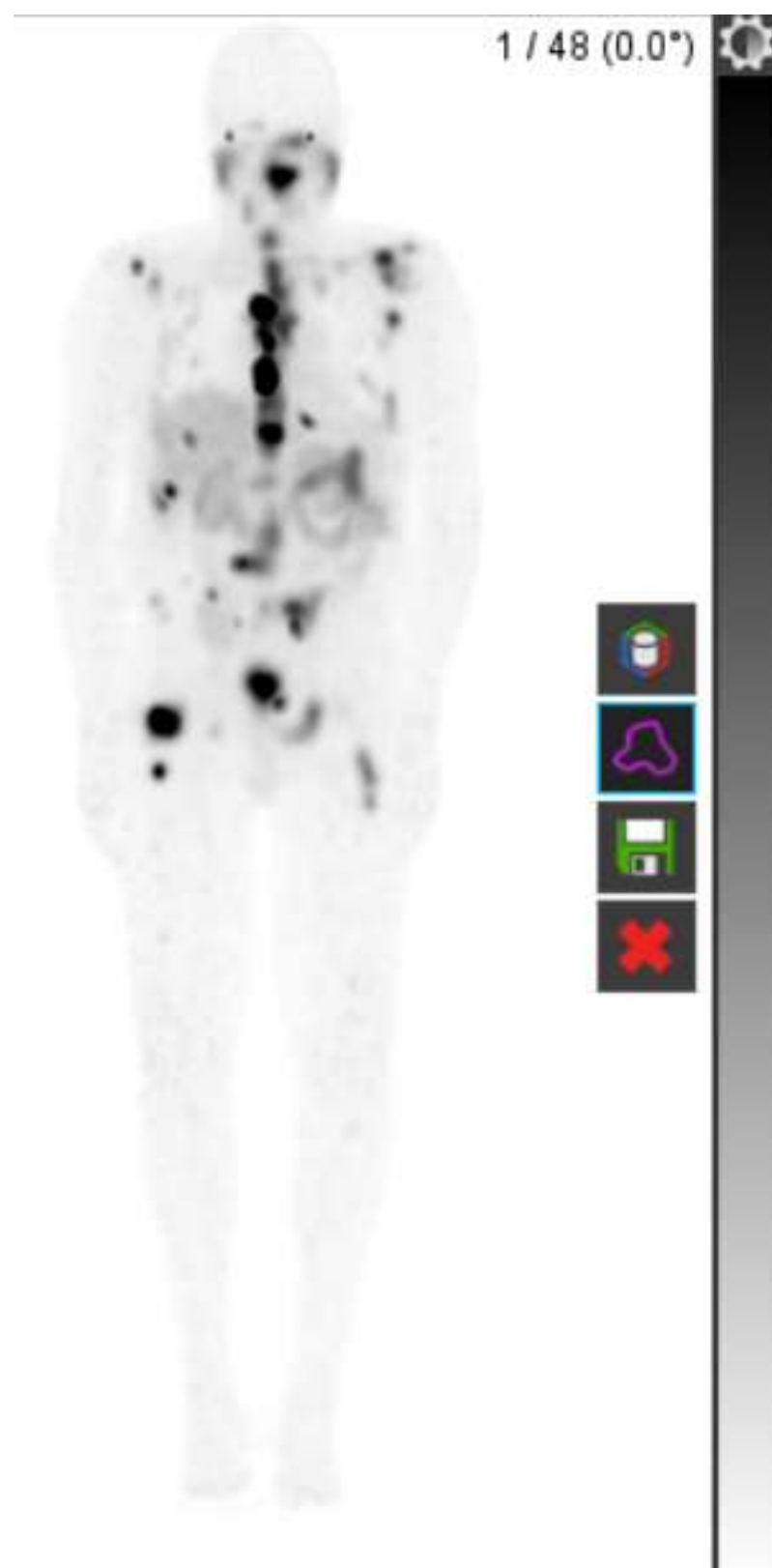




LU-177 SPECT/CT IMAGING

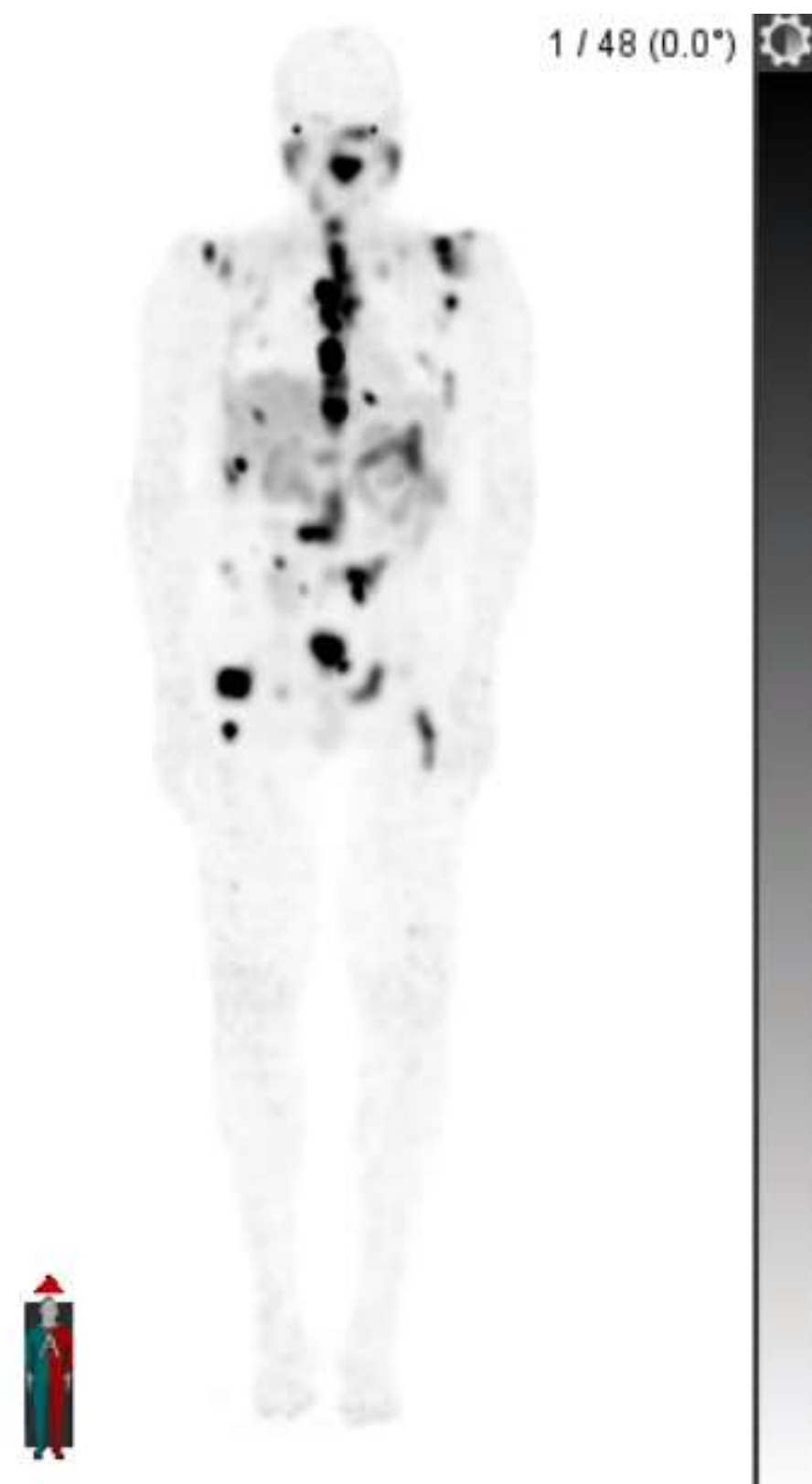
Veriton 200 Series

113 keV peak



18 min

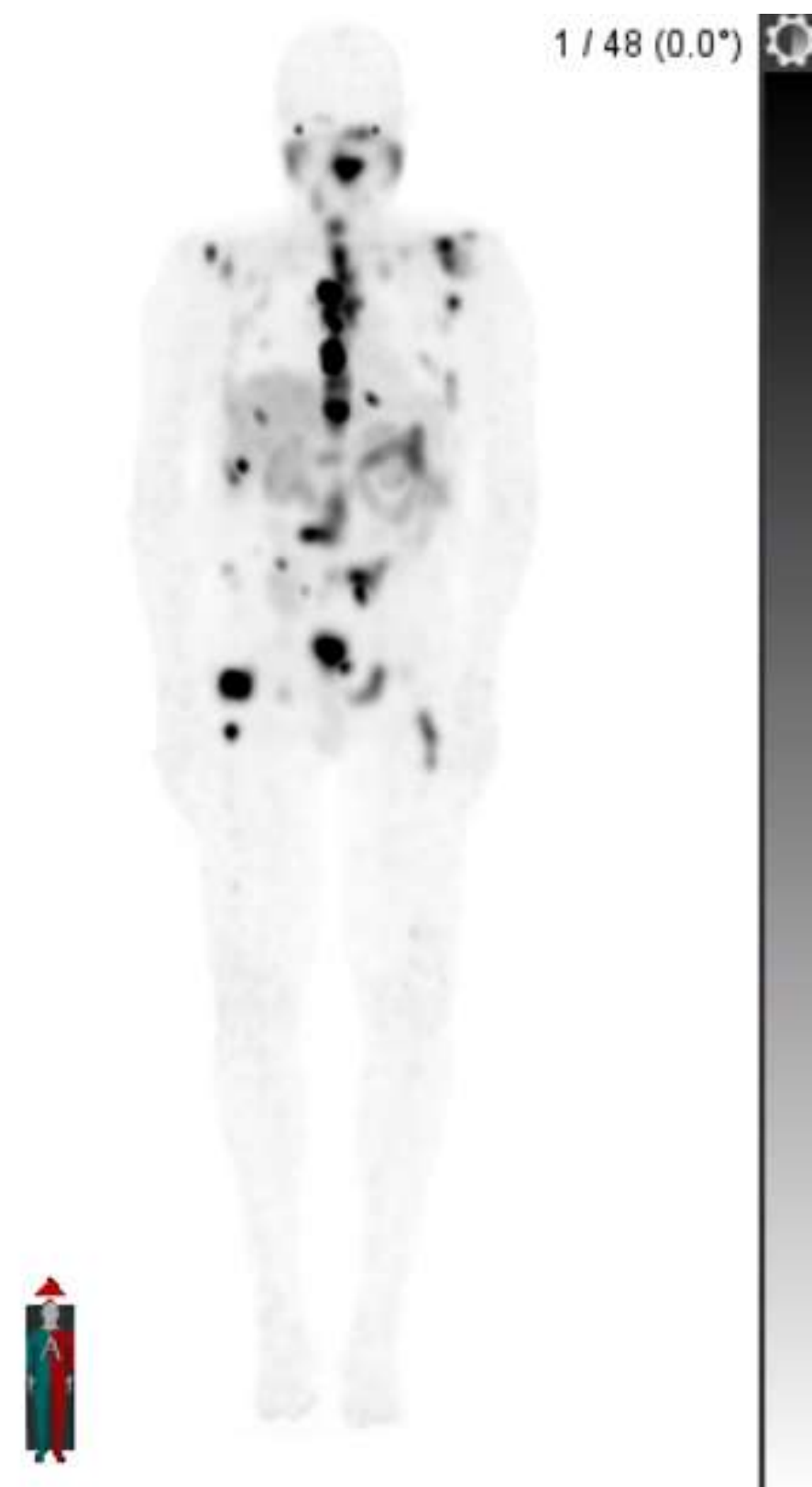
208 keV



18 min

Veriton 400 Series

113 & 208 keV

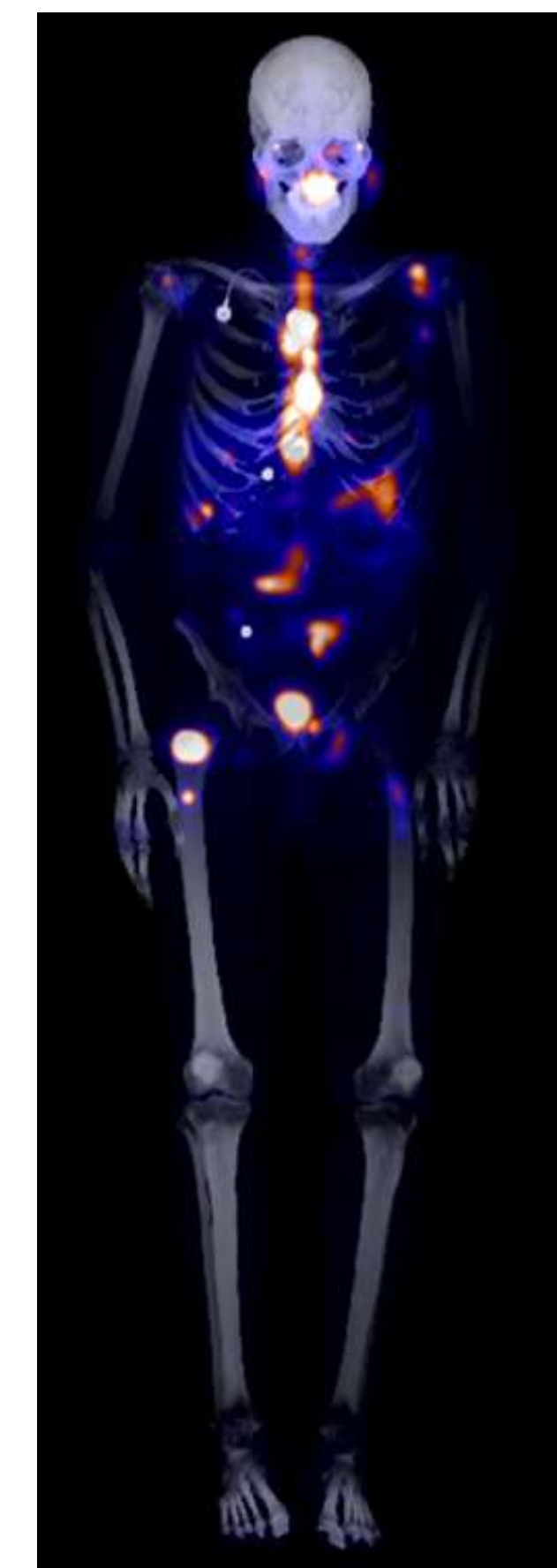


18 min

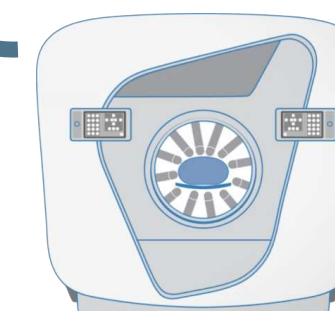
113 & 208 keV

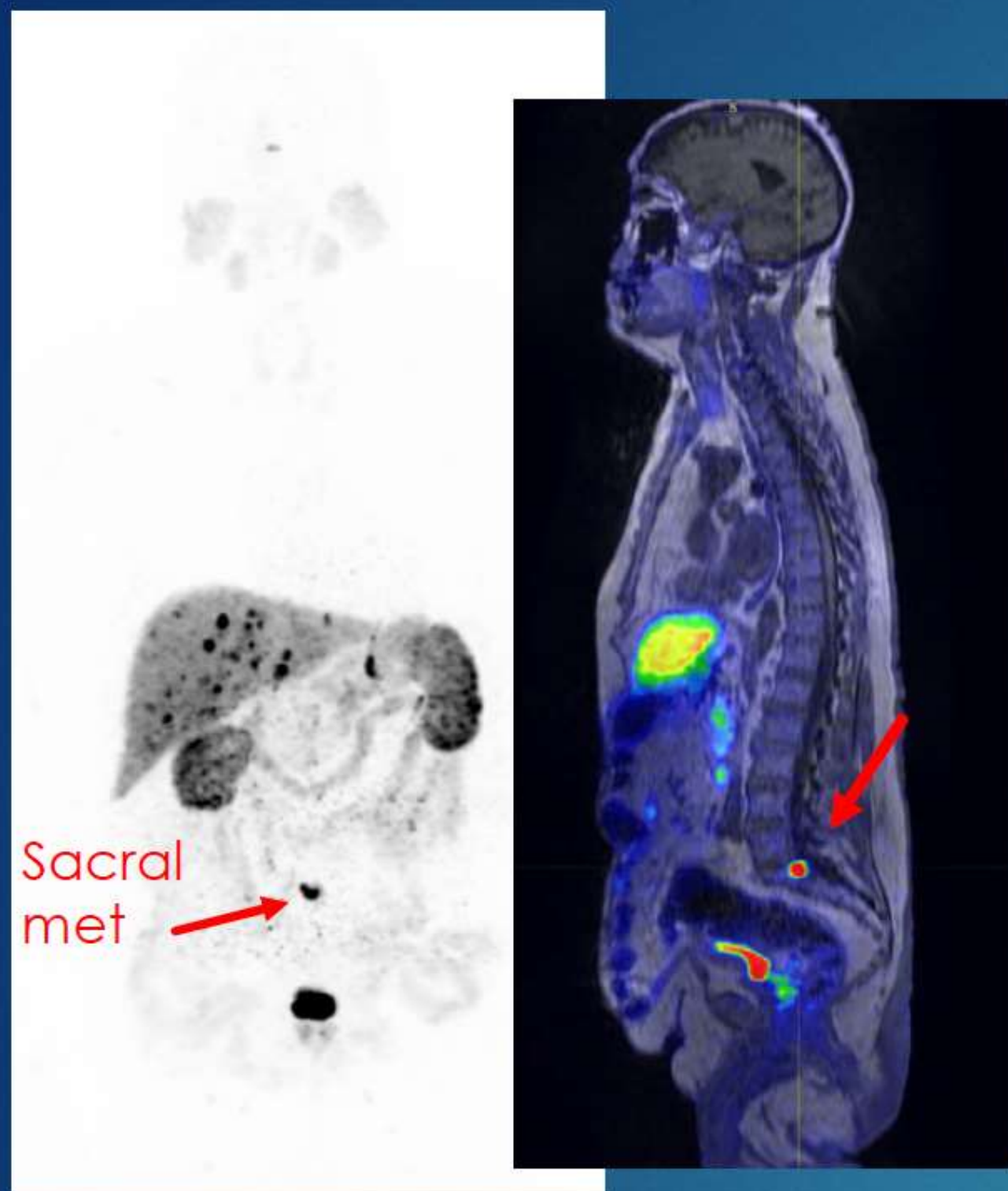


9 min!



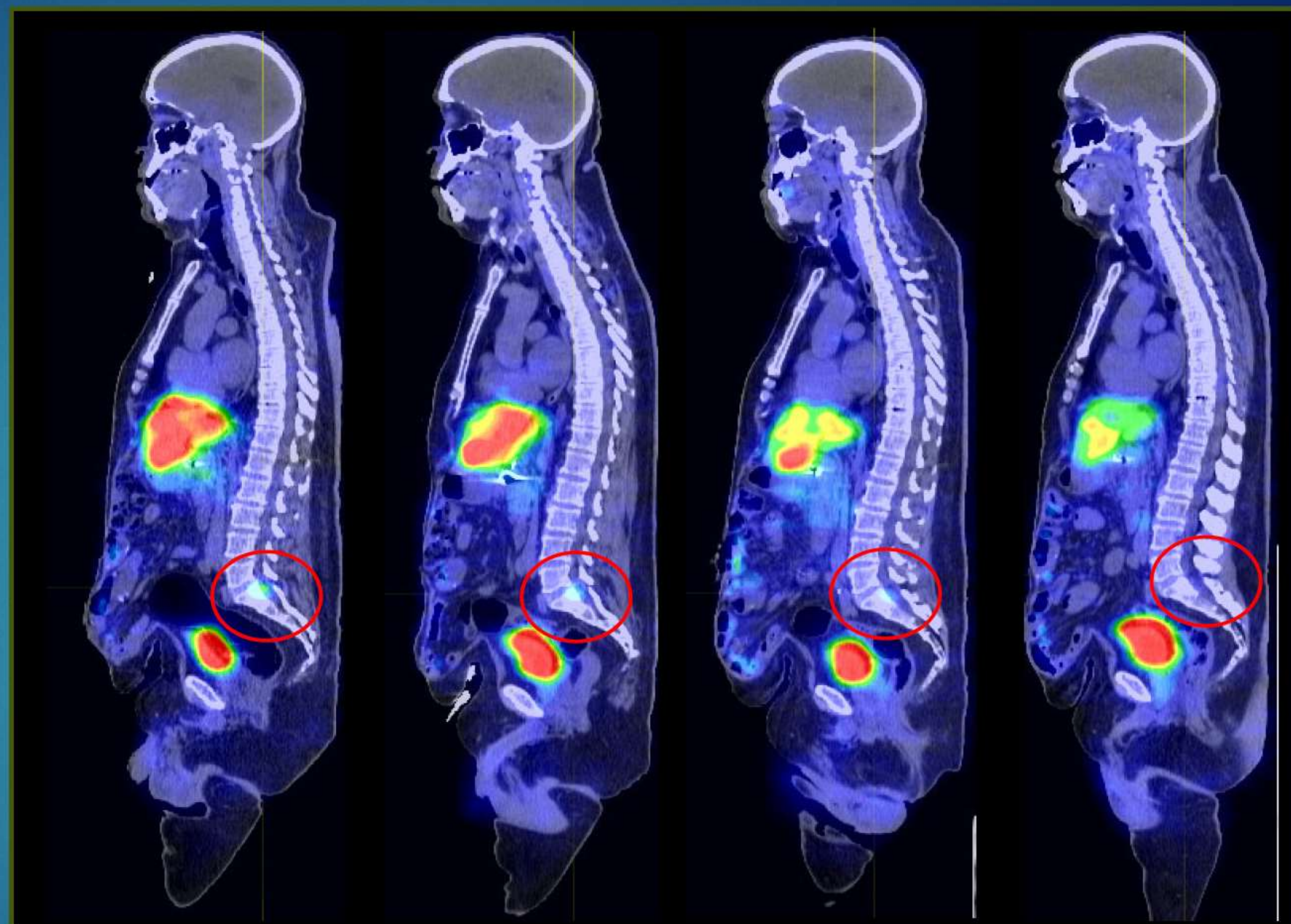
SPECT/CT Fusion





Ga-68 Dotatate
PET/MR Survey

8 mo prior to Lu-177
therapy



Cycle 1
Max SUVbw = 2.9

Cycle 2
1.9

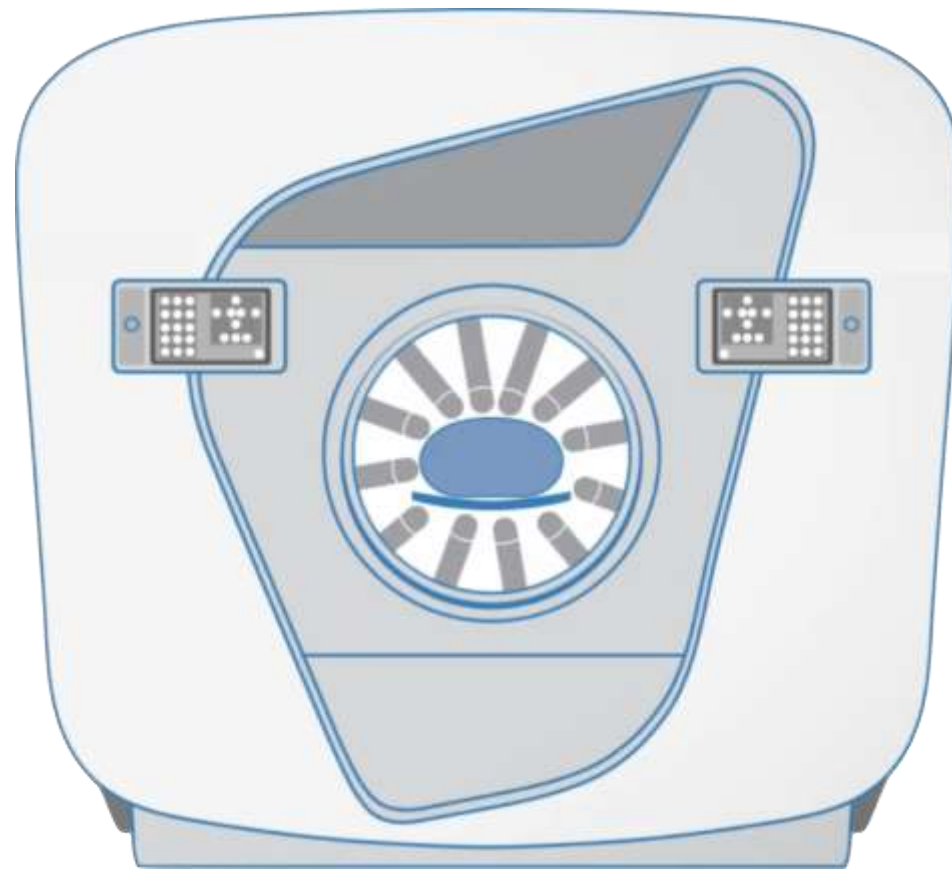
Cycle 3
1.5

Cycle 4
1.0

4-6 hrs after Lutathera administration, imaged on Veriton SPECT/CT

Images courtesy of the
Mayo Clinic, Rochester, MN, USA

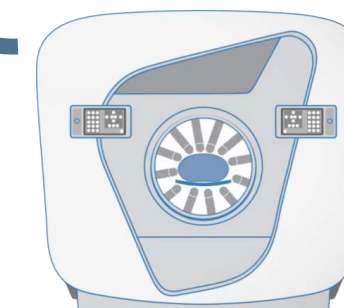
Absolute Quantitation



CT Attenuation and Scatter Correction
 Resolution Recovery
 Sophisticated Reconstruction Algorithm

Image natively produced in Activity Concentration (Bq/ml)

→ **Need for Correction Factor!**

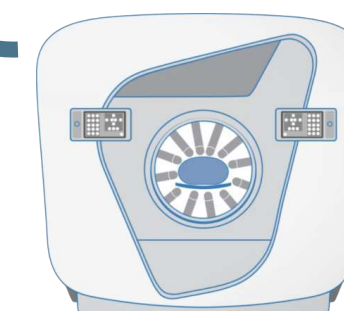




WHAT to Calibrate? — Image Counts Into Activity Concentration (Bq/ml)

Controllable technical factors related to image acquisition and formation?

Parameters	Conventional SPECT	Full-Ring CZT SPECT	PET
Angular sampling	✓	✗	✗
Acquisition Range (180/360)	✓	✗	✗
Orbit type and distance	✓	✓	✗
Collimators	✓	✗	✗
Matrix Size / Voxel Size	✓	✓	✓
Acquired Counts	✓	✓	✓
Nuclide / Energy Window	✓	✓	✗
Scatter Correction Method	✓	✓	✗
Reconstruction Parameters (# iteration, filters, algorithm,...)	✓	✓	✓





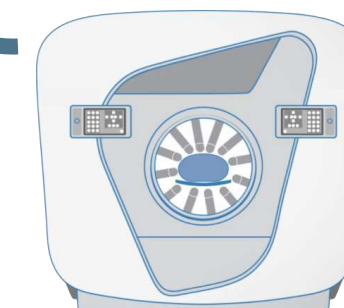
HOW to Calibrate?

Cross-Calibration against Dose-Calibrator with Uniform Flood Phantom

Parameters	Full-Ring CZT SPECT
Angular sampling	✗
Acquisition Range (180/360)	✗
Orbit type and distance	✓
Collimators	✗
Matrix Size / Voxel Size	✓
Acquired Counts	✓
Nuclide / Energy Window	✓
Scatter Correction Method	✓
Reconstruction Parameters (# iteration, filters, algorithm,...)	✓

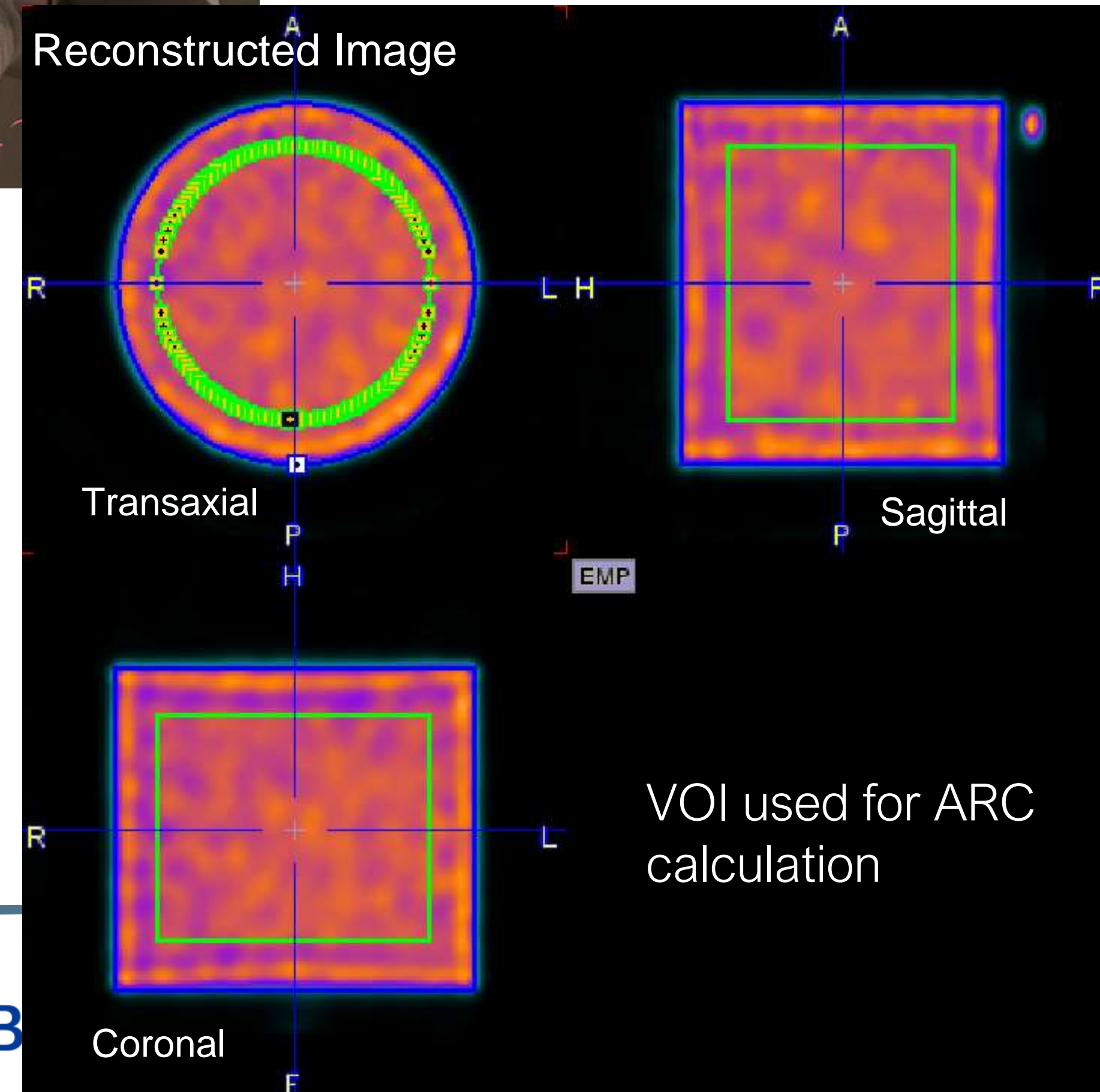
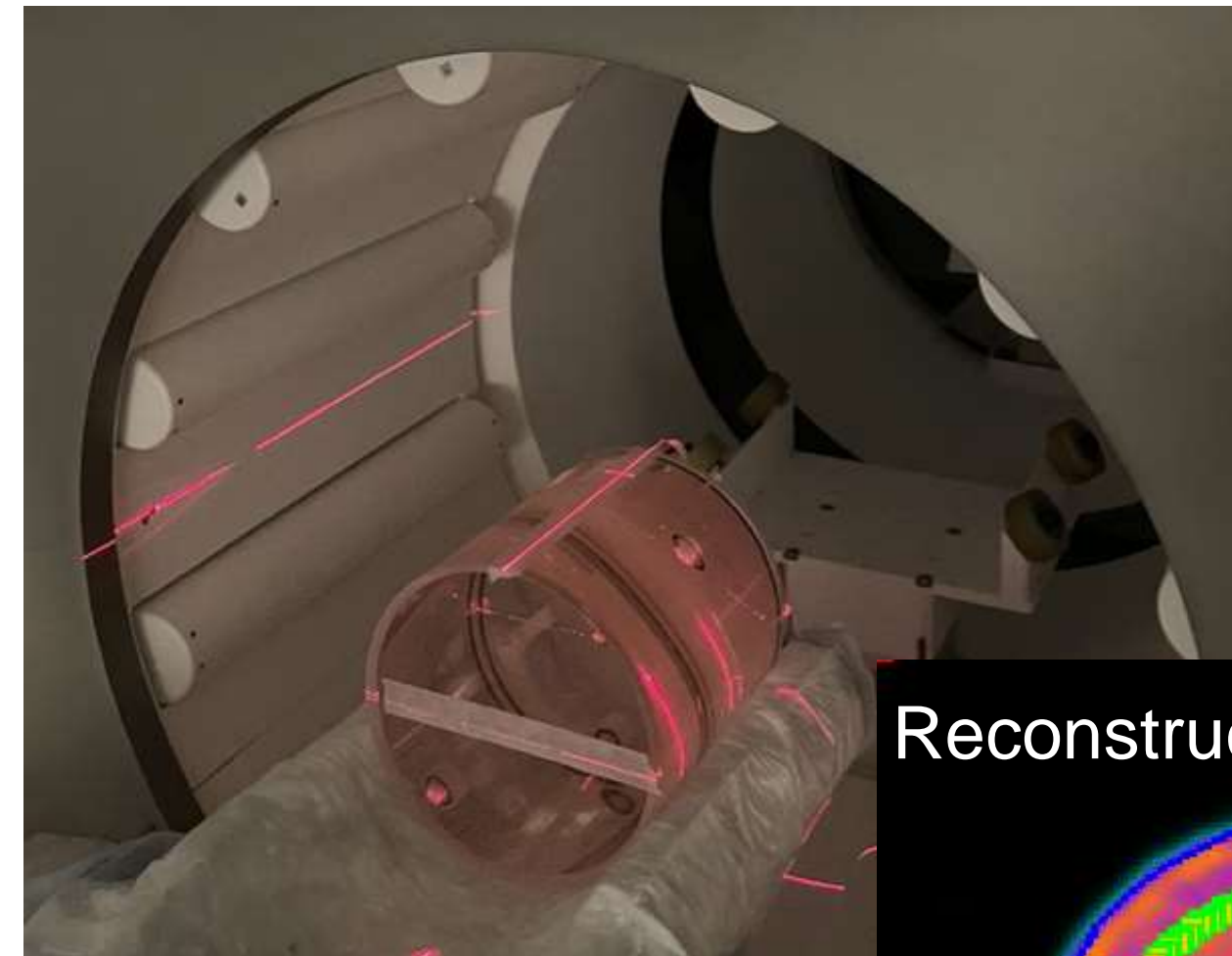


Uniform Flood Cylinder (as in PET)
 Calibrated against Dose Calibrator
 How Robust are these factors? Variability?



Uniform Flood Calibration

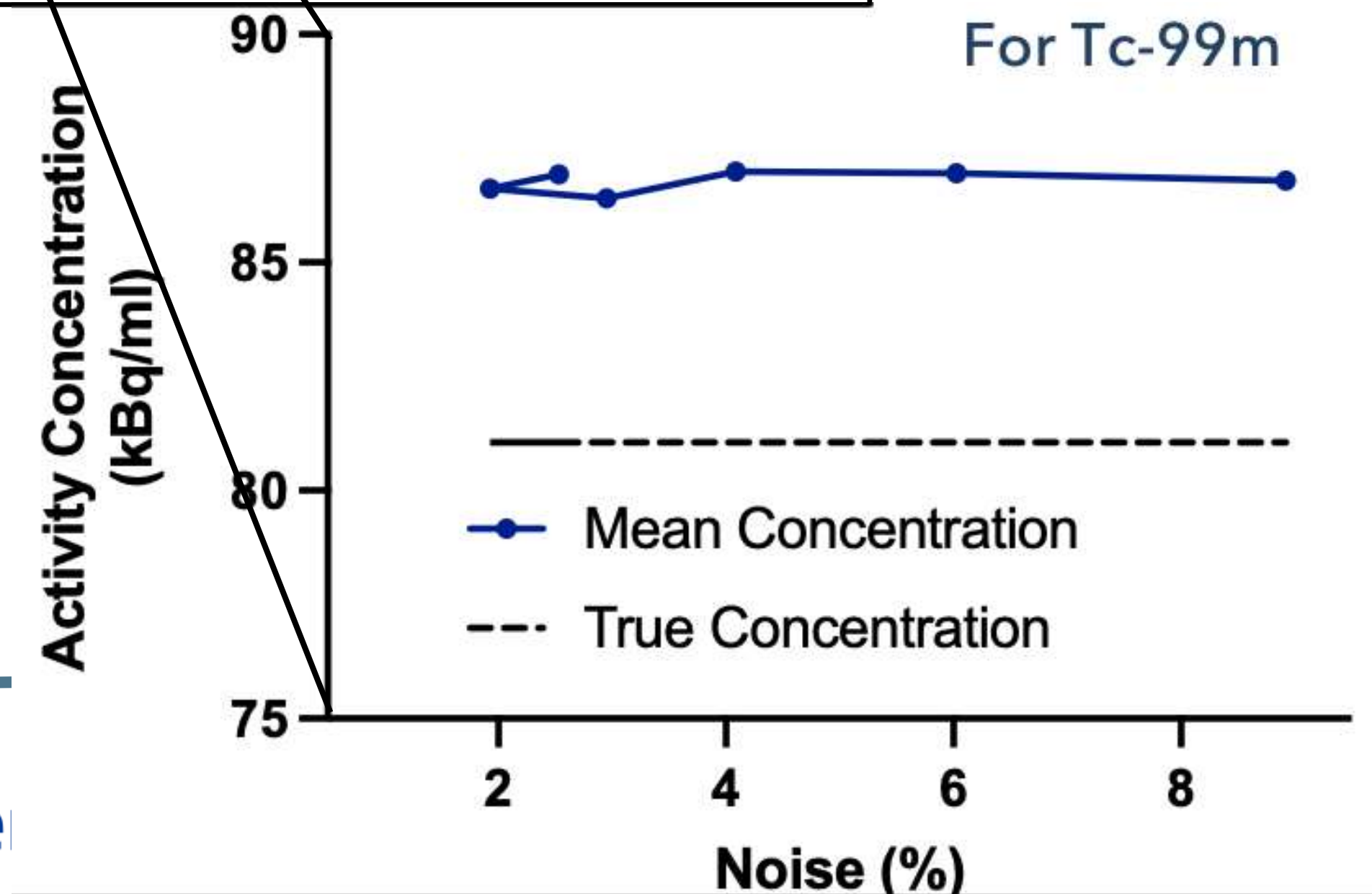
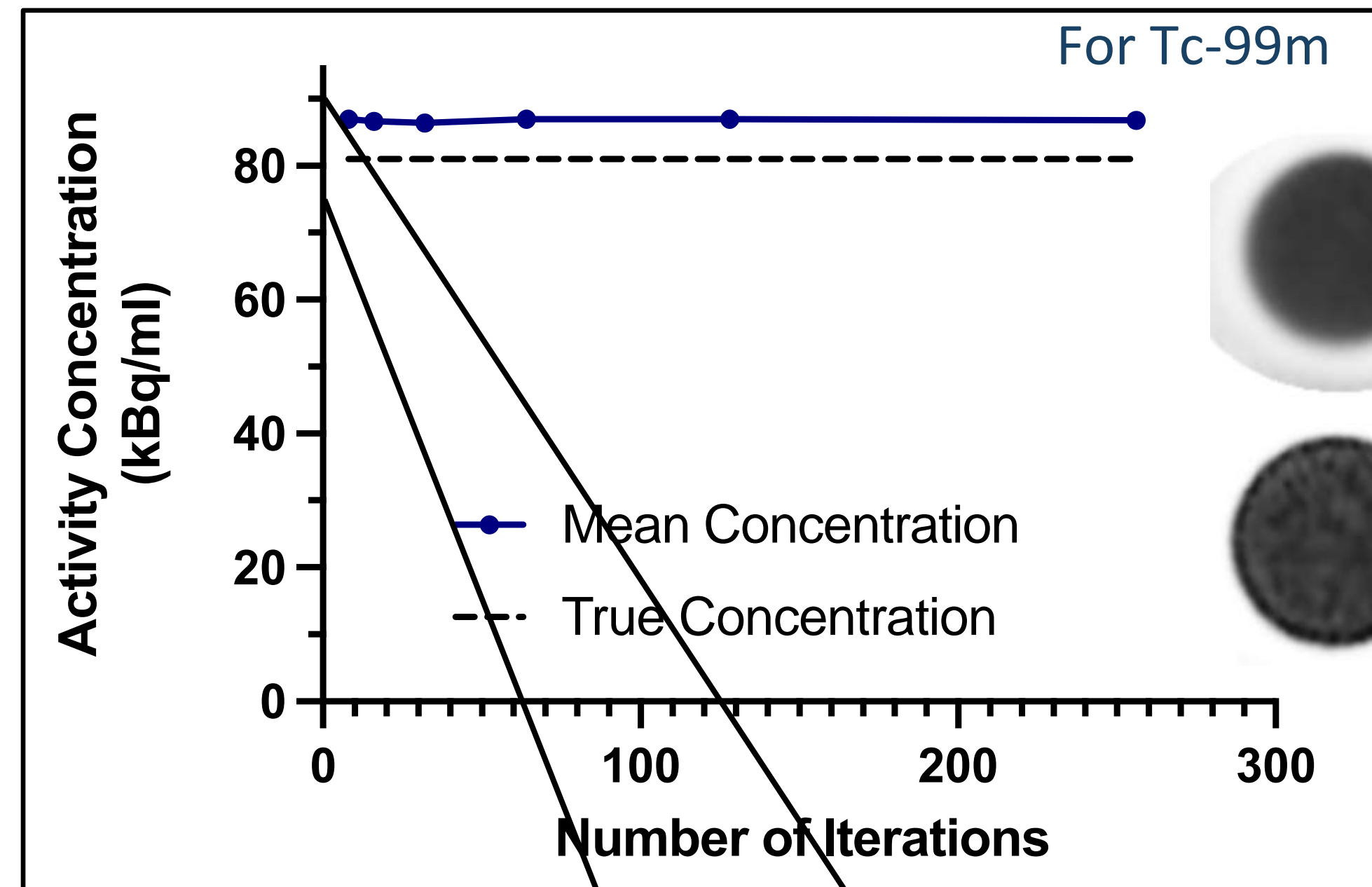
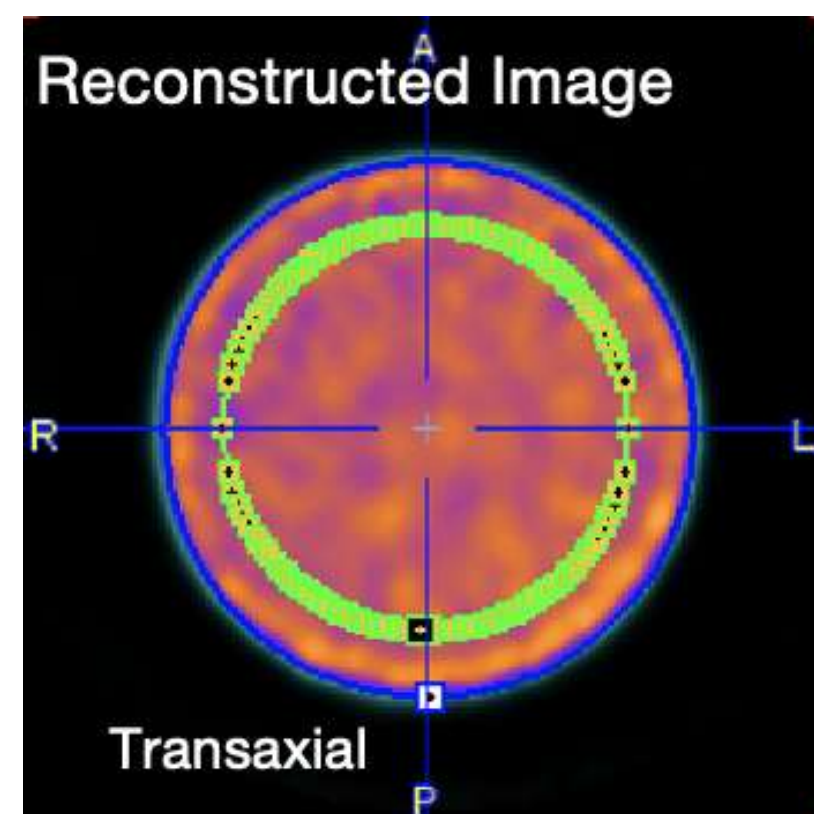
- Acquisition of a **Uniform Flood phantom** with **clinical protocol parameters**
- High Count level (32 Mcts)** to minimize noise
- Reconstruction parameters and scatter correction** similar to **clinical protocol parameters**
- Estimation of the **Activity Recovery Coefficient ARC** or **Calibration Factor** ($= \text{image concentration} / \text{true concentration}$).
True concentration obtained from known activity (DC) and phantom volume (6415 ml)



How Many Iterations is Optimal?

- ARC is stable over iterations
- Not too many to avoid incorporating **excessive noise** (*increases variability*)
- Not too few to ensure convergence

VOI used for ARC calculation

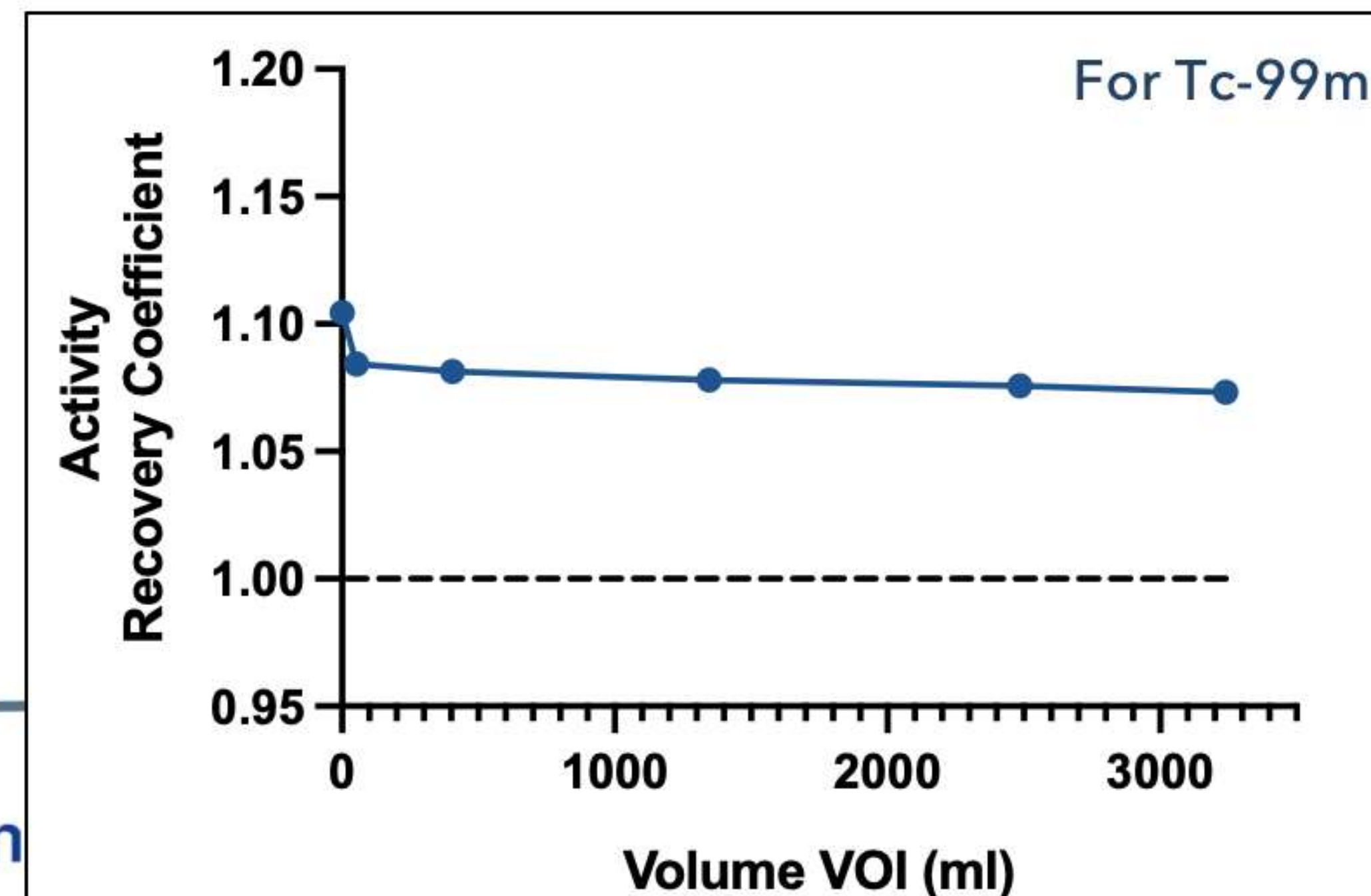
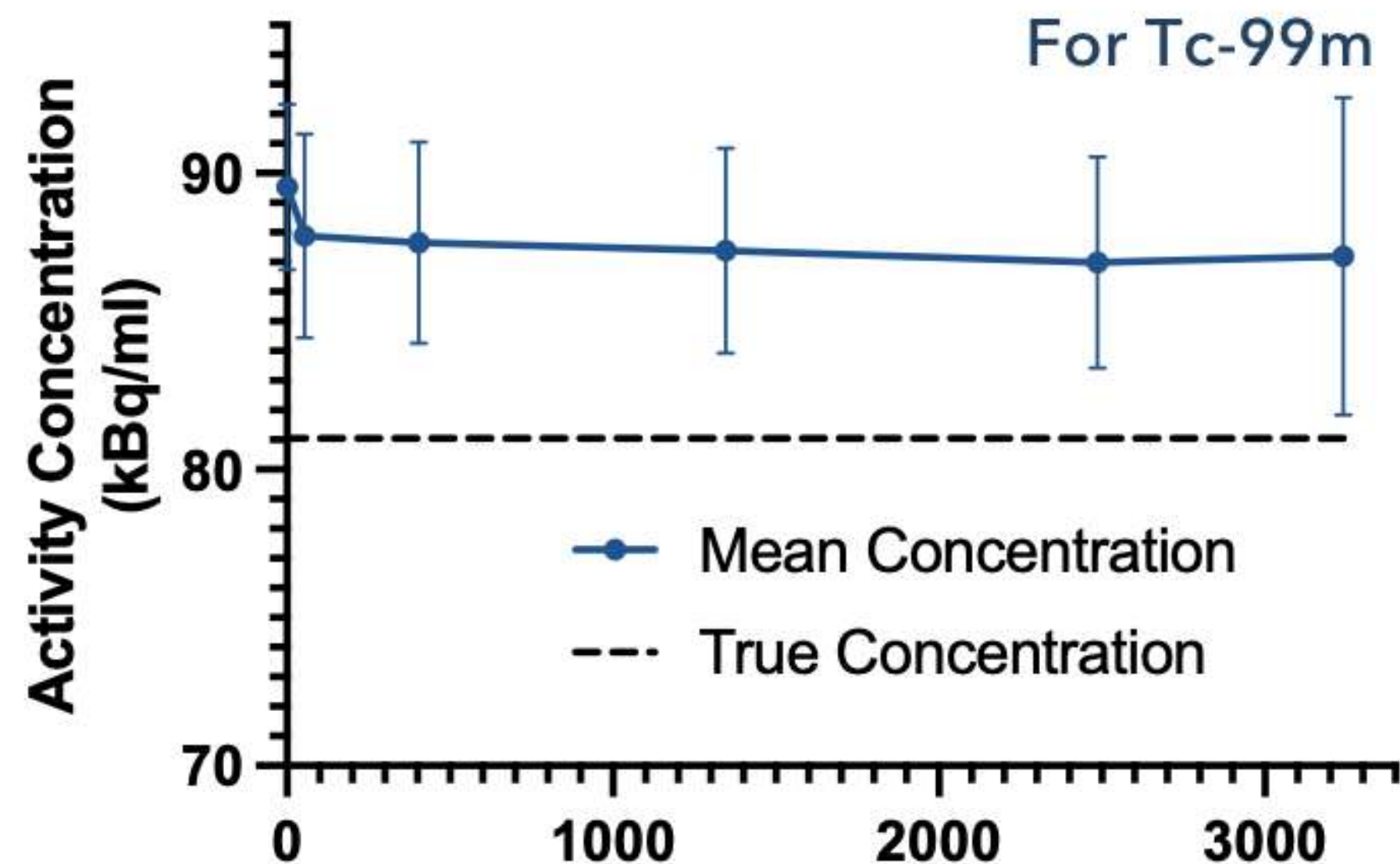
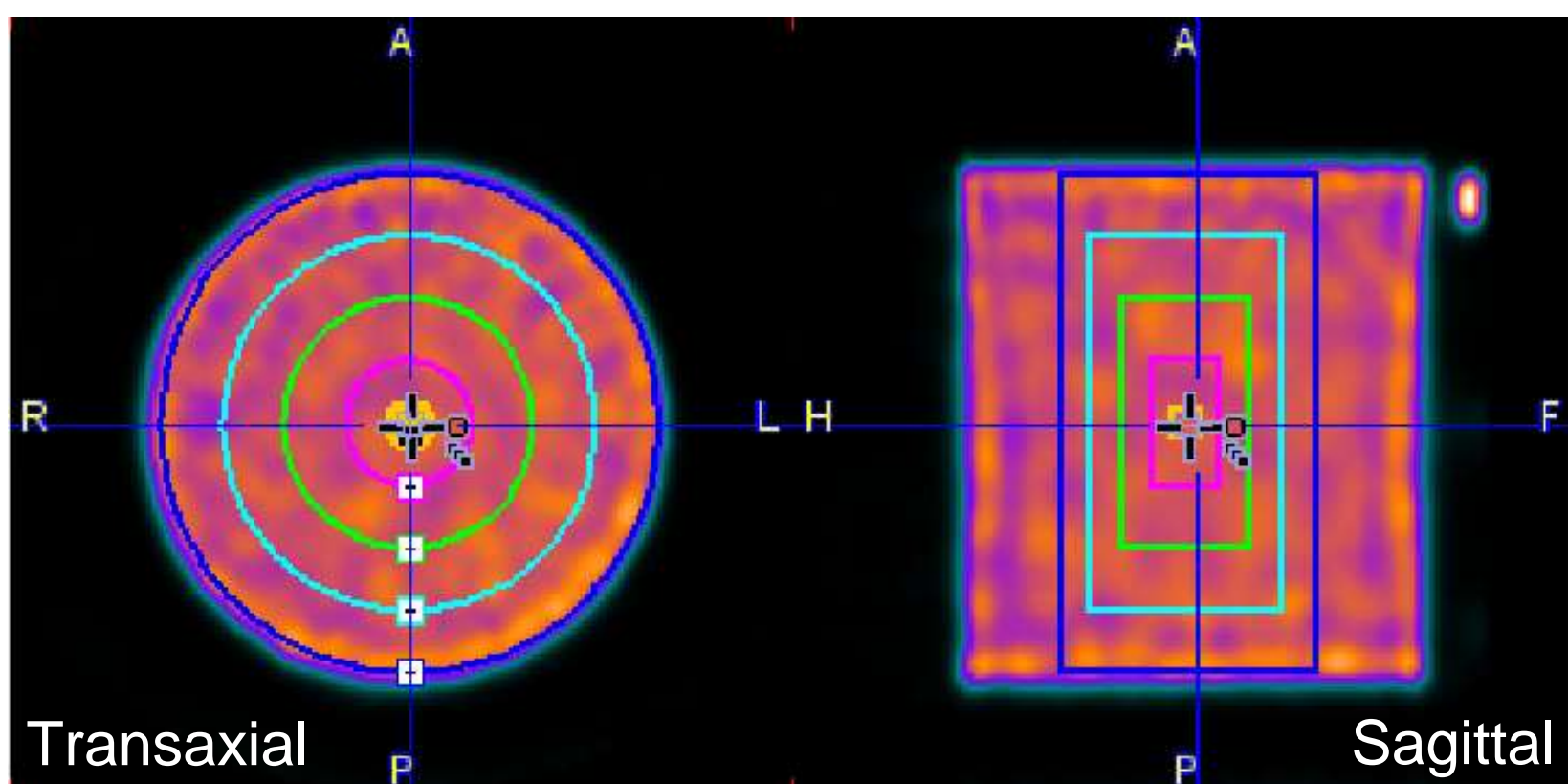


What Should be the VOI size?

Not too small to avoid incorporating excessive noise (*increases variability*). Higher likelihood of including more scatter/attenuation effects in small VOIs.

Not too large to exclude edge effects

VOI used for ARC calculation

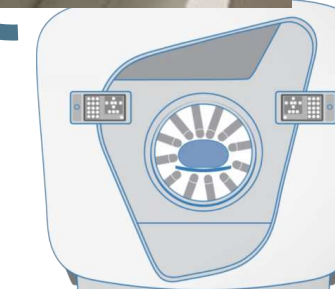
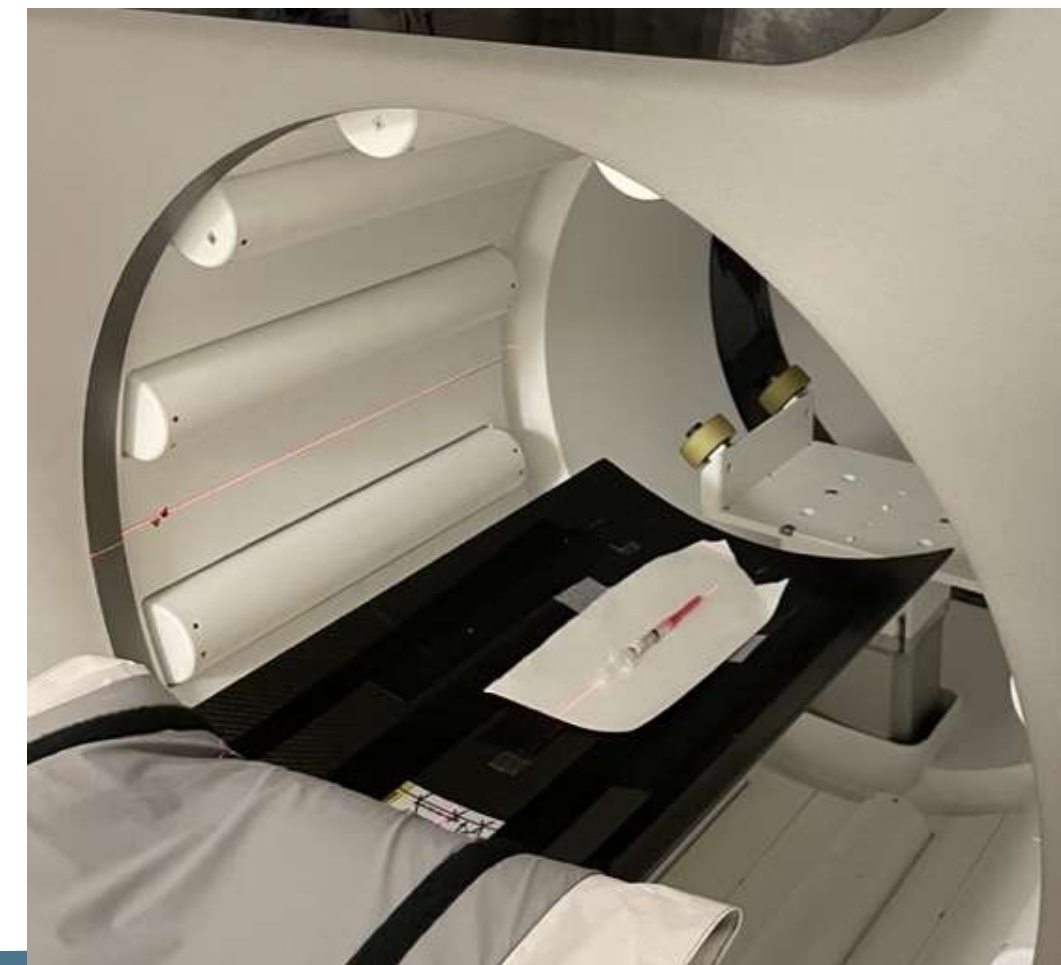


Can any other source objects be used for calibration?

Uniform Flood phantom might not be adequate for long-lived nuclides (e.g., Lu-177, In-111, I-131) in practice as it needs to be stored until it has decayed.

Flask or **Syringe** would be more convenient in practice.

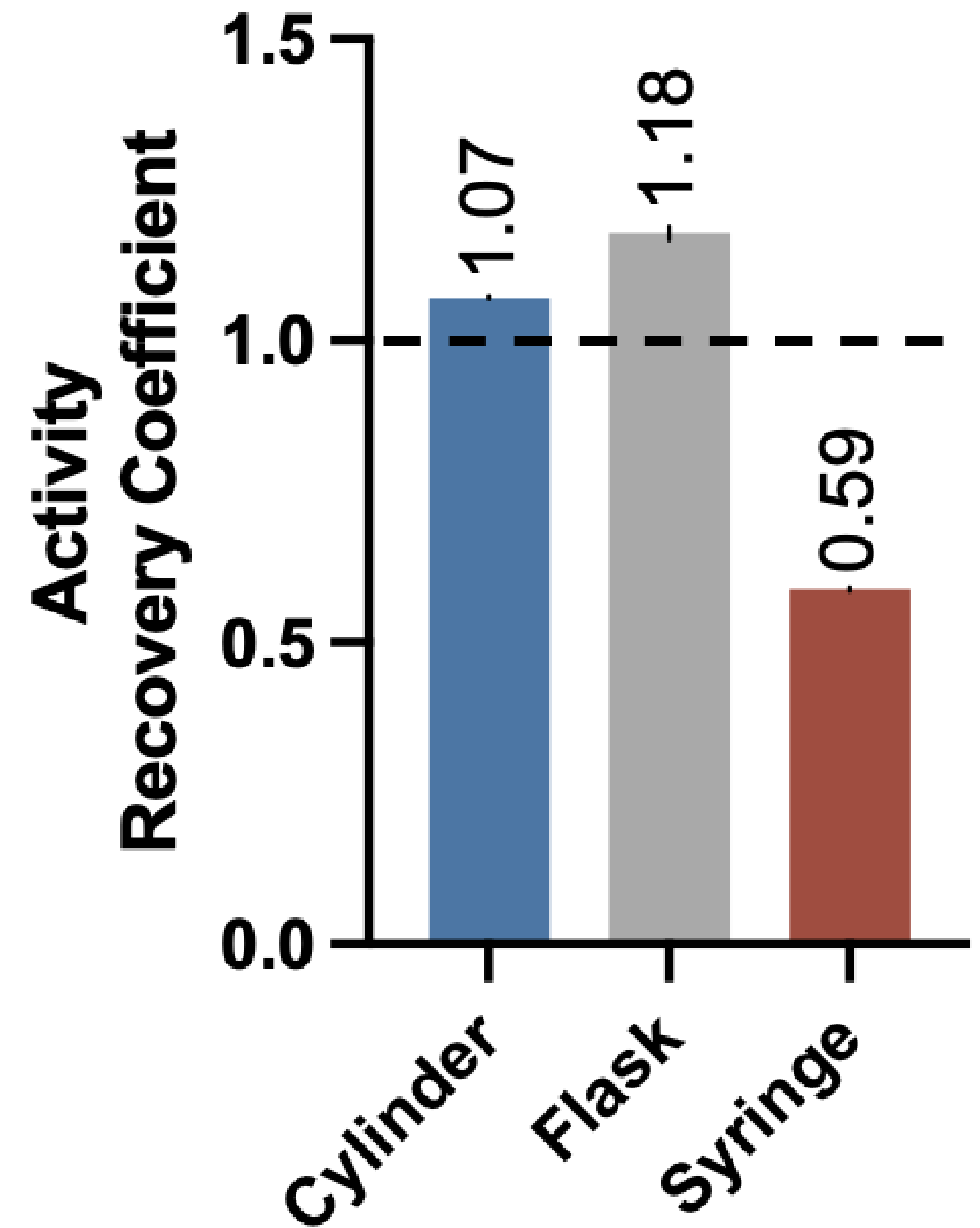
What is the impact of the calibration source objects on absolute quantitation?



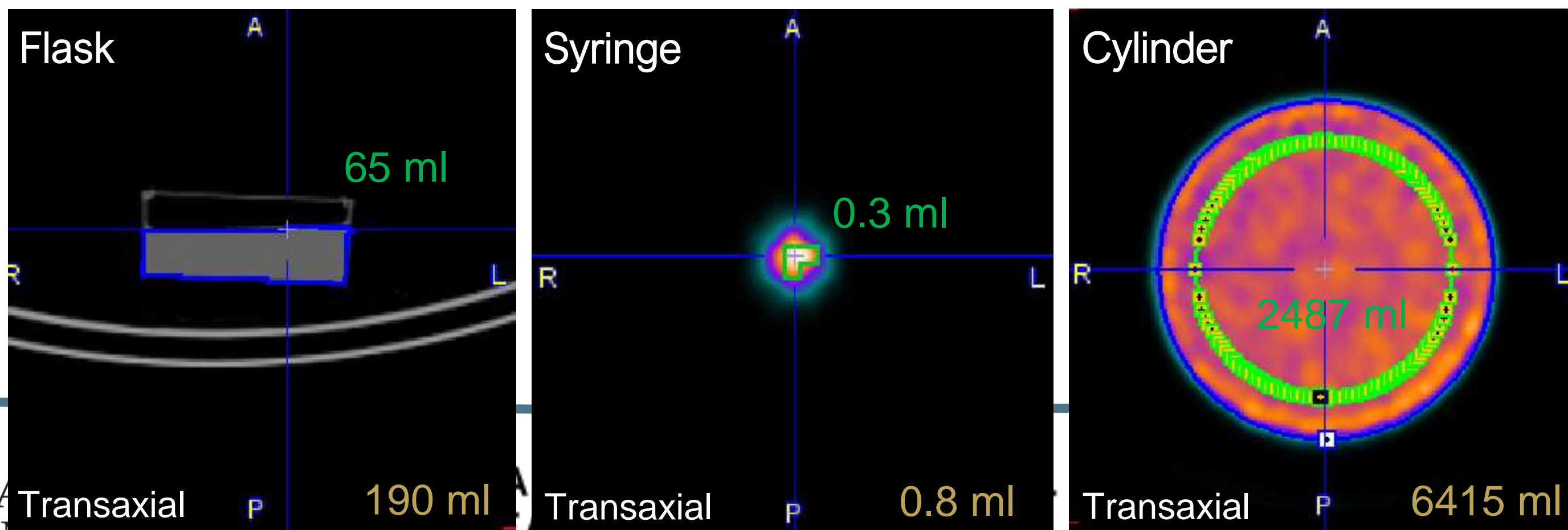
Can any other source objects be used for calibration?

ARC is **overestimated** with the flask (*too many counts detected*) likely due to Gibbs edge artefact (*Resolution Recovery*) [1].
 Using this calibration factor will lead to **under-correction**.

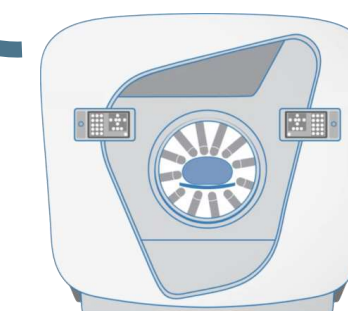
ARC is **underestimated** with syringe (*too few counts recorded*) due to partial volume effect (spill over).
 Using this calibration factor will **over-correct**.



VOI used for ARC calculation



[1] Armstrong *Nuc. Med. Commun.* 2016



What is the Impact of the Calibration Accuracy on Absolute Quantitation?

Cardiac Torso Phantom ^{99m}Tc-MPI Study

Clinical Count level (stress): 1.6-3 Mcts

True Activity Concentration

Myocardium: 543.3 kBq/ml

Blood Pool: 61.9 kBq/ml

Liver: 148.8 kBq/ml

Background: 49.6 kBq/ml

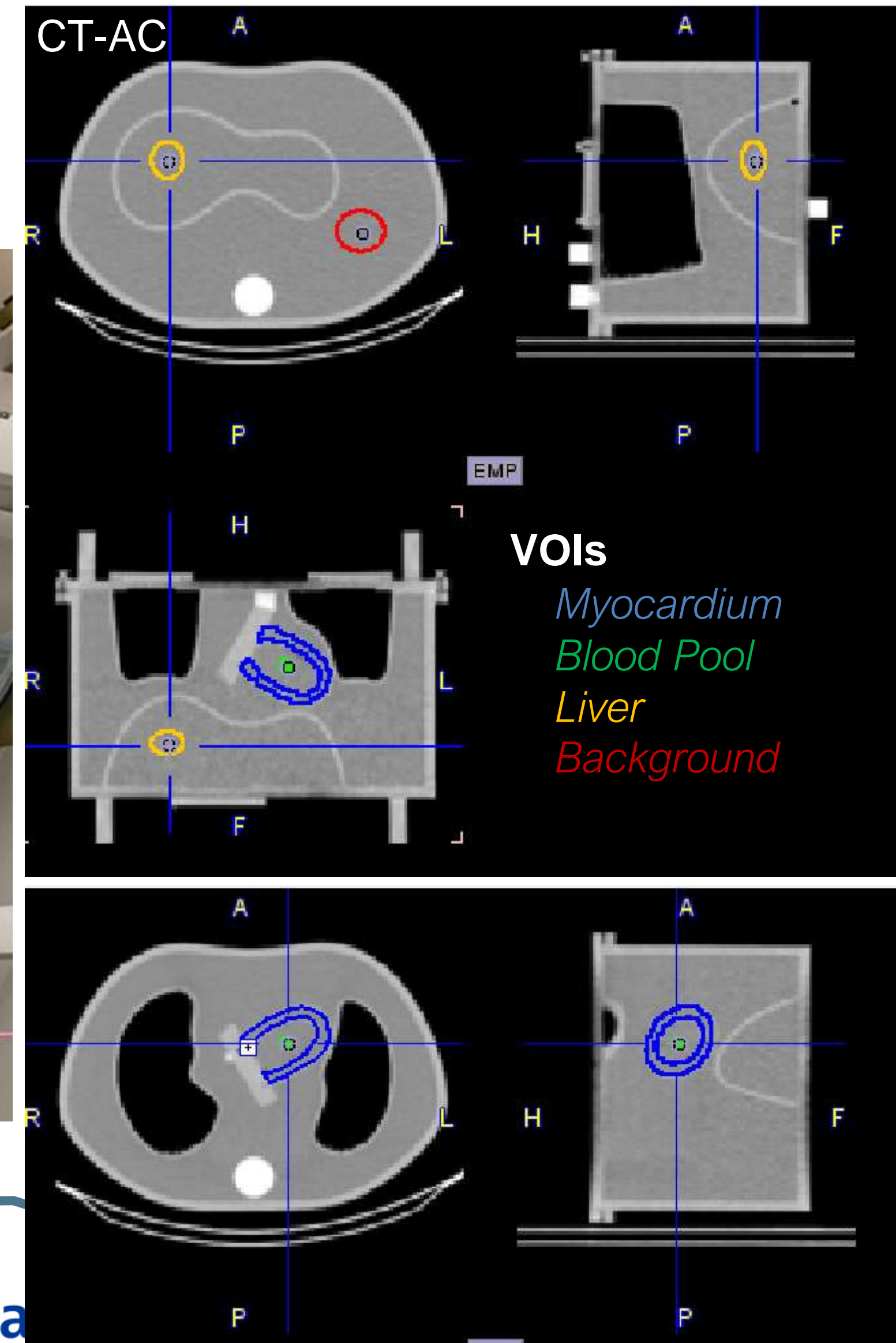
VOIs derived from CT-AC (for Myocardium)

Myocardium: 99.6 ml

Blood Pool: 0.5 ml

Liver: 13.2 ml

Background: 43 ml

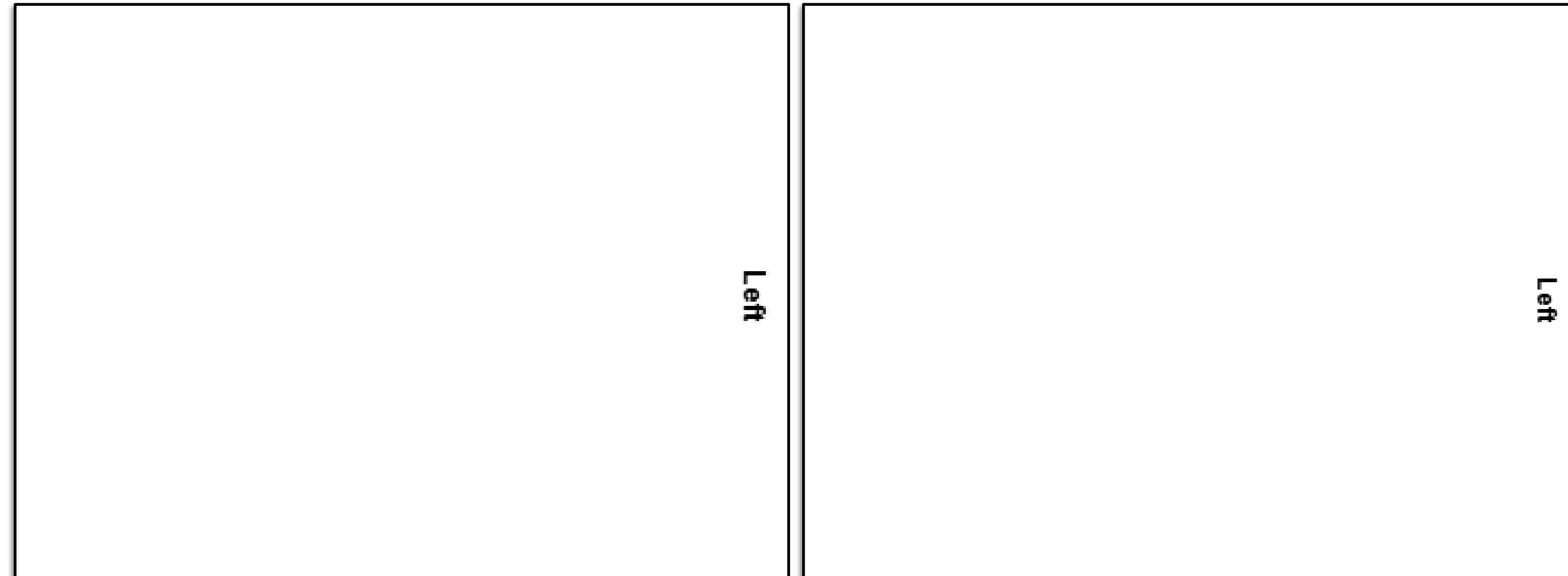


What is the Impact of the Calibration Accuracy on Absolute Quantitation?

Cardiac Torso Phantom Study
 Clinical Count levels (Stress ^{99m}Tc-MPI)
 4 it. 8 sub. OSEM + RR/AC/SC

1.6 Mcts (Transaxial Fly Through)

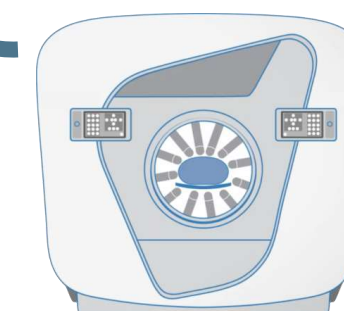
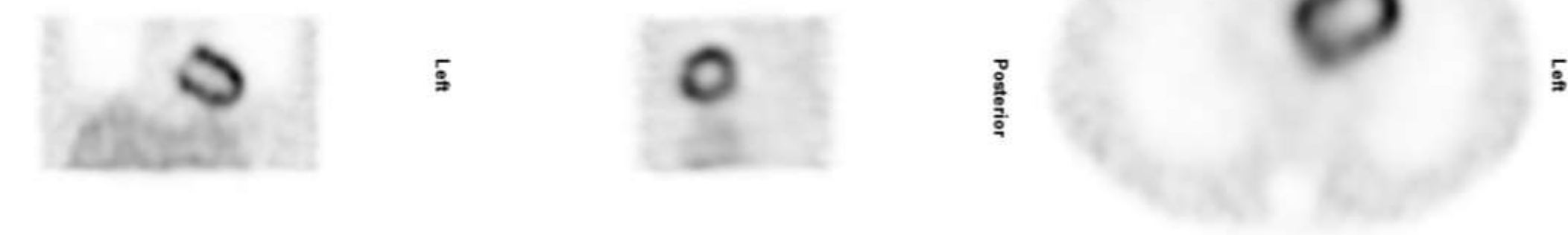
3 Mcts (Transaxial Fly Through)



1.6 Mcts



3 Mcts



What is the Impact of the Calibration Accuracy on Absolute Quantitation?

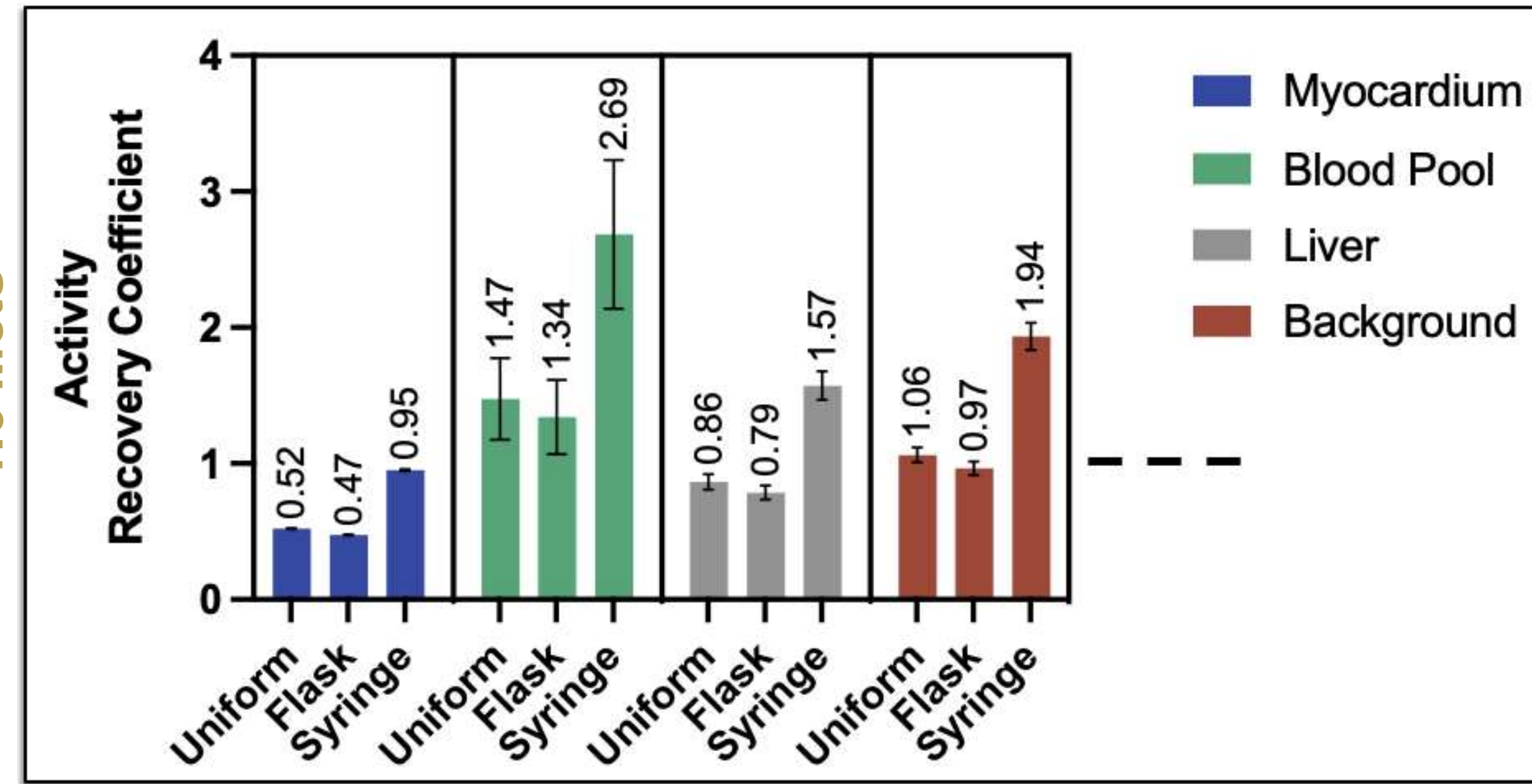
Cardiac Torso Phantom Study
 Clinical Count levels (Stress ^{99m}Tc-MPI)
 4 it. 8 sub. OSEM + RR/AC/SC

Calibration derived from the **uniform cylinder** leads to the **most accurate quantitation** (ARC~1) on overall.

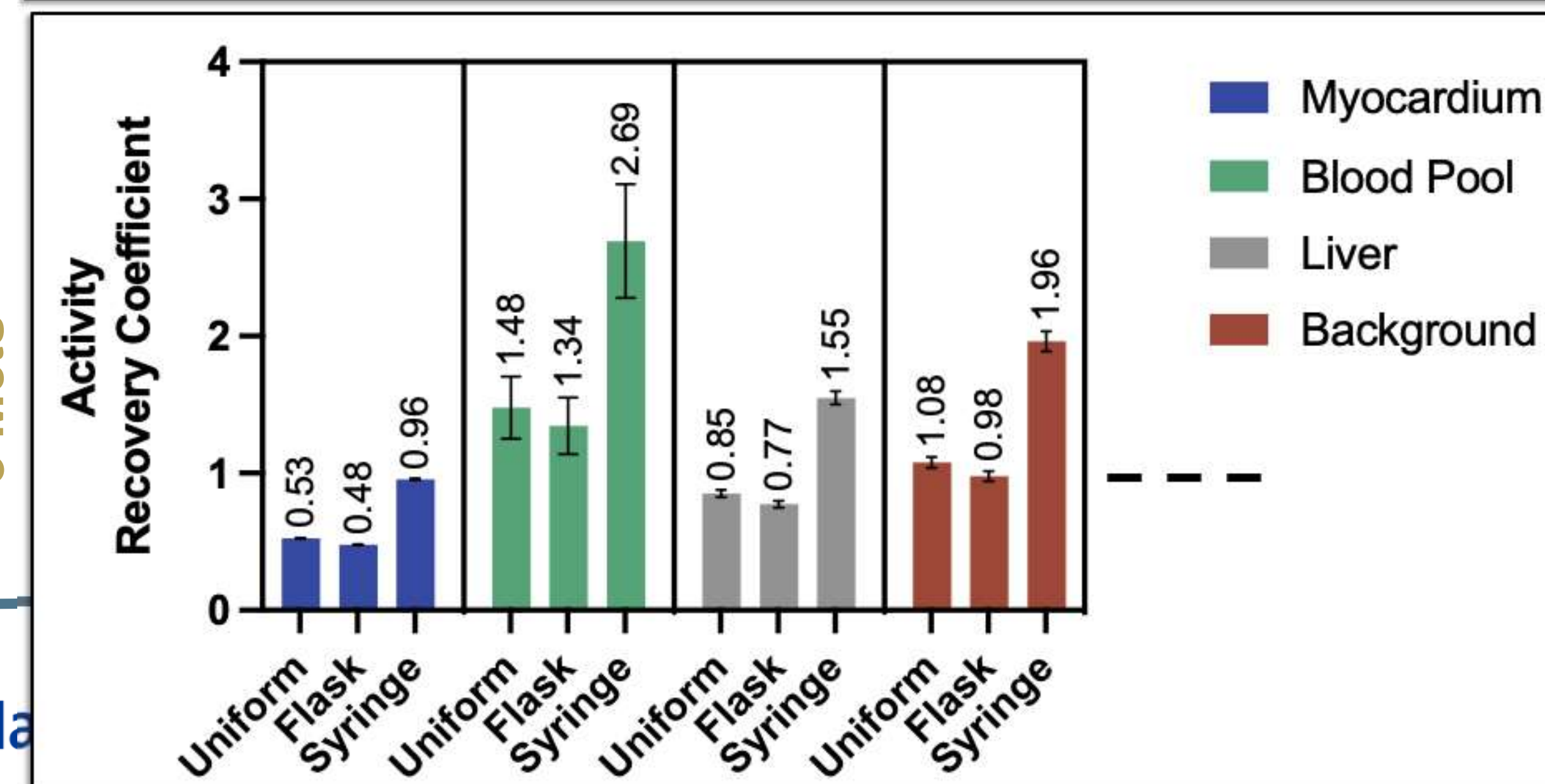
Flask Calibration underestimates overall the true concentration, achieving the **lowest ARCs**.

Syringe Calibration strongly **overestimates true concentration**. However, by chance, the syringe calibration led to the best **ARC for the myocardium**, as the syringe and the myocardium were **similarly affected** by PVE.

1.6 Mcts



3 Mcts



Absolute Calibration derived from a Jaszczak Phantom Acquisition?

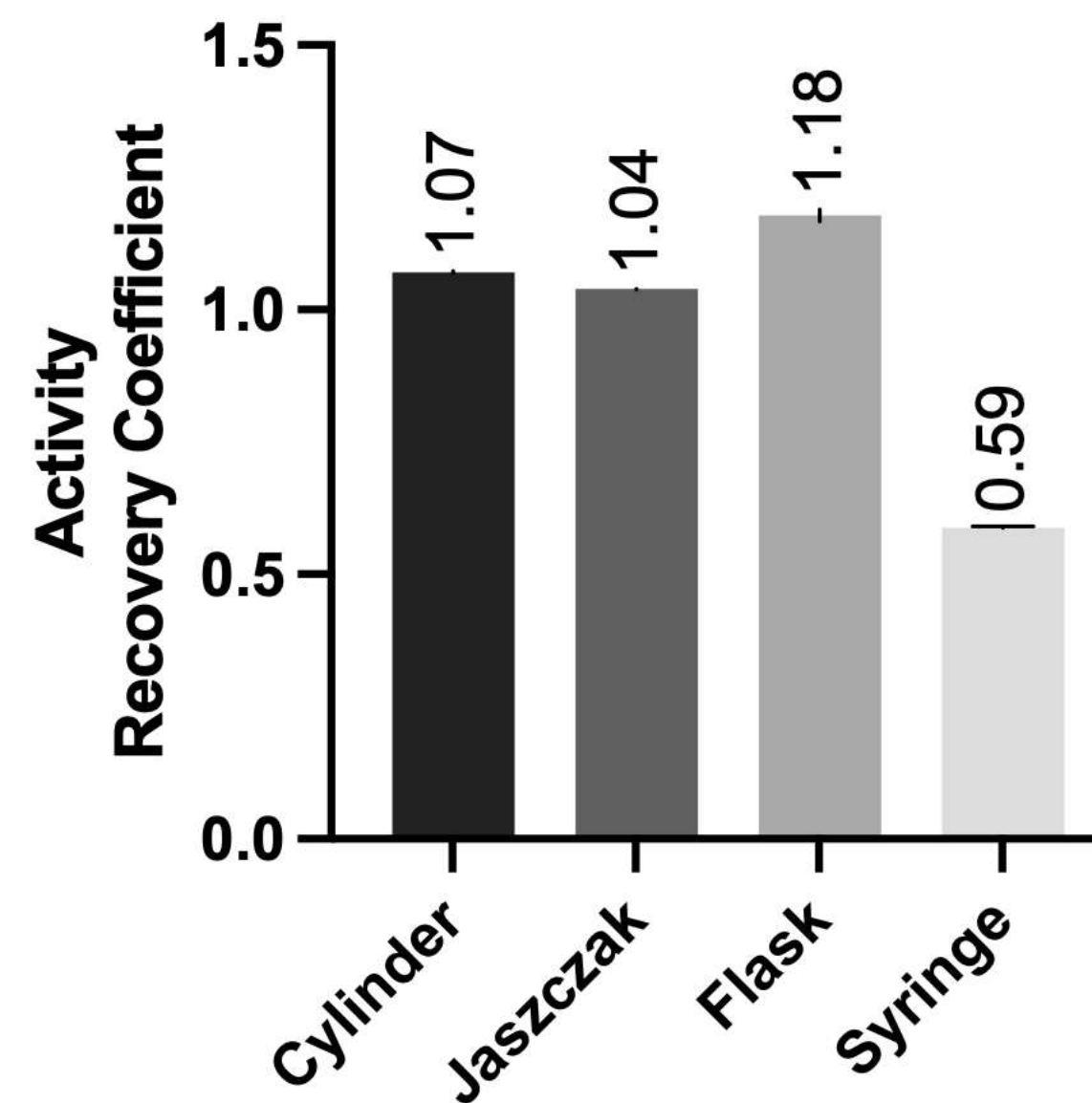
Calibration Scan

10 Mcts Non-Focus Mode (^{99m}Tc)
4 it. 32 sub. OSEM + RR/AC/SC

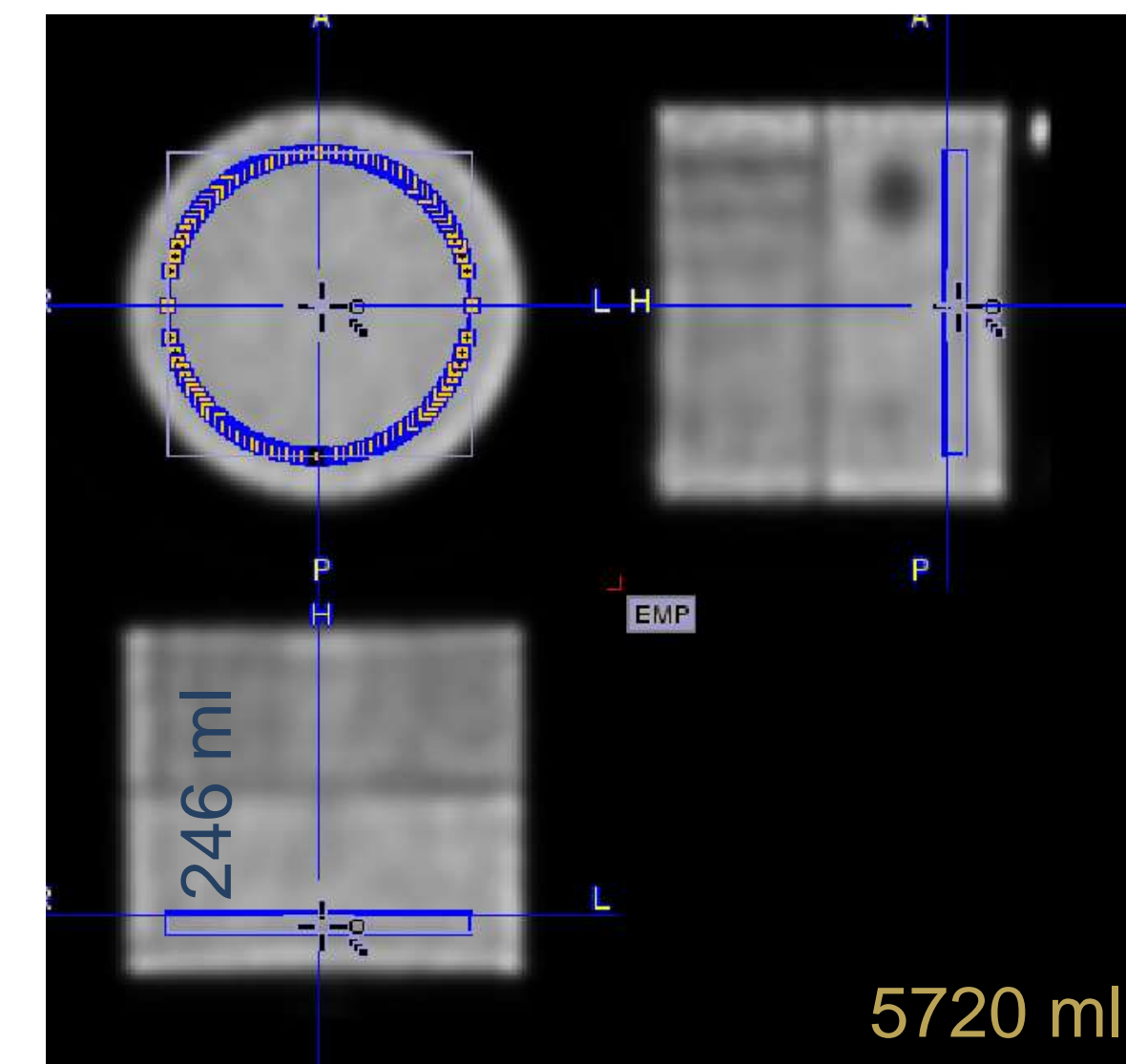
Jaszczak phantom acquisition (*usually*) enters quarterly QC.

Easier to derive a calibration factor from such frequent scans than with uniform phantom!

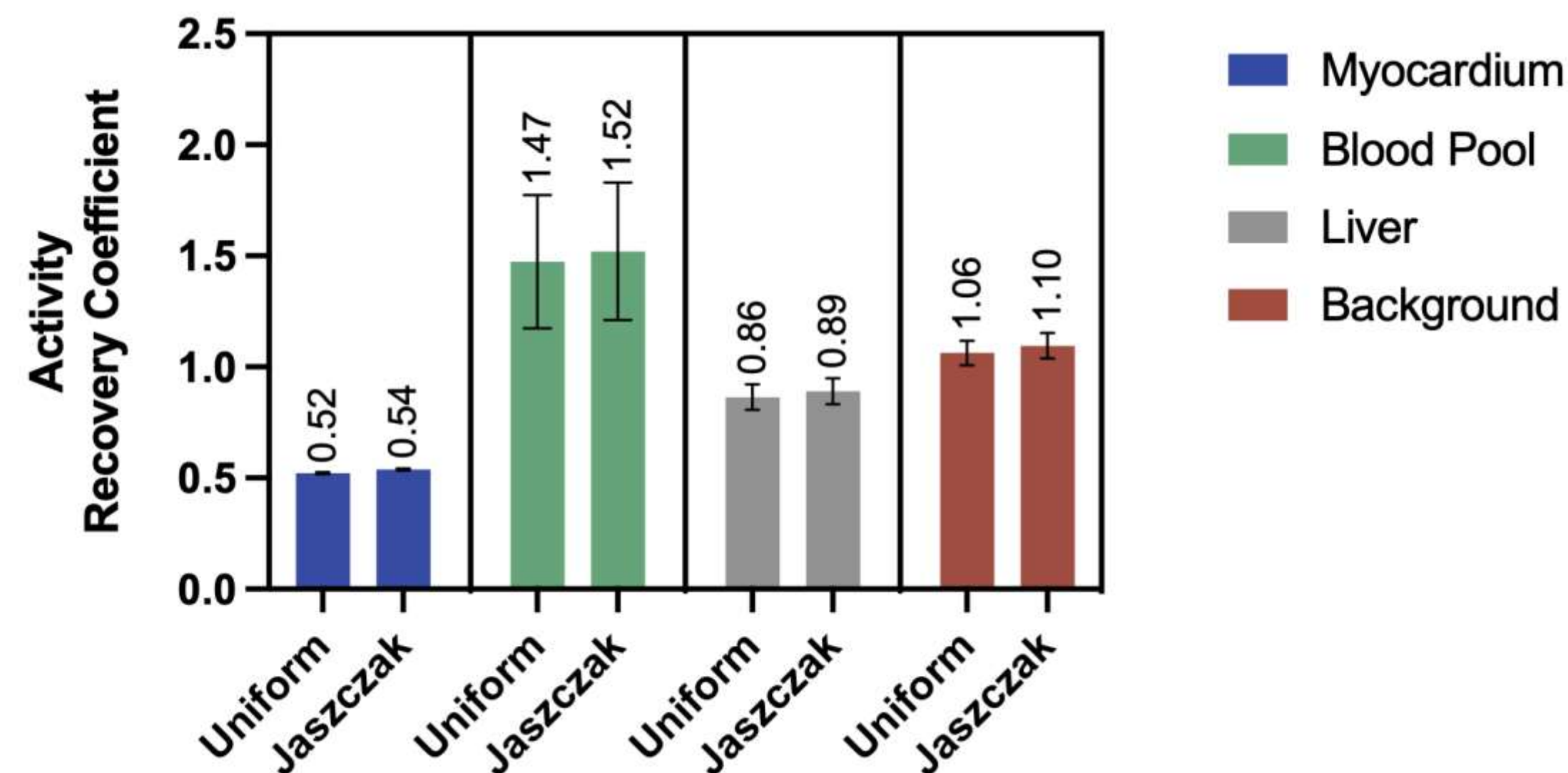
Calibration factor remains close to what is derived from a uniform cylinder.
Small Impact on Absolute Quantitation (*assessed torso phantom*)!



VOI used for ARC calculation



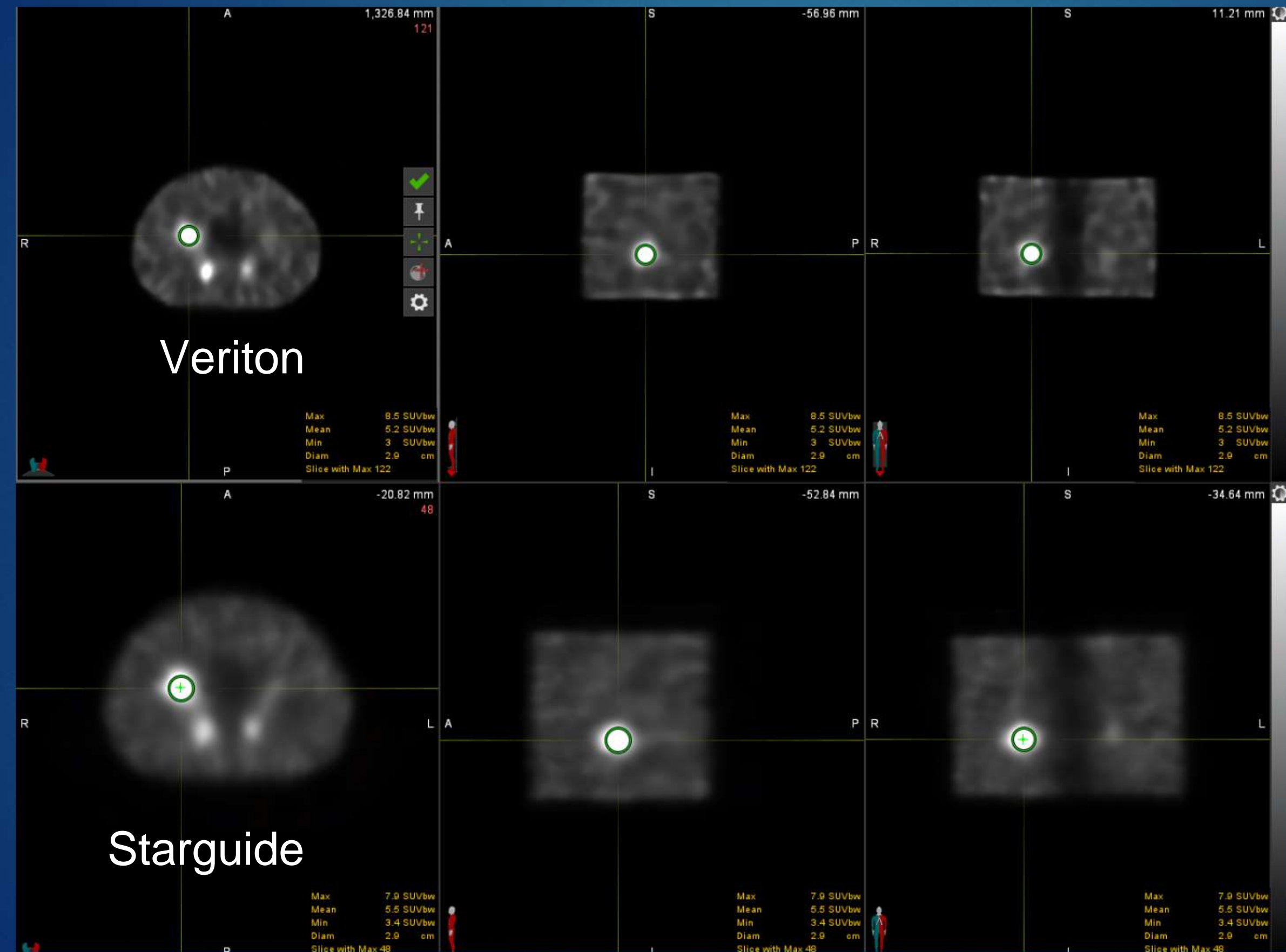
Absolute Quantitation Jaszczak Vs Uniform





QUANTITATION: VERITON/STARGUIDE

Veriton or StarGuide? Pretty close

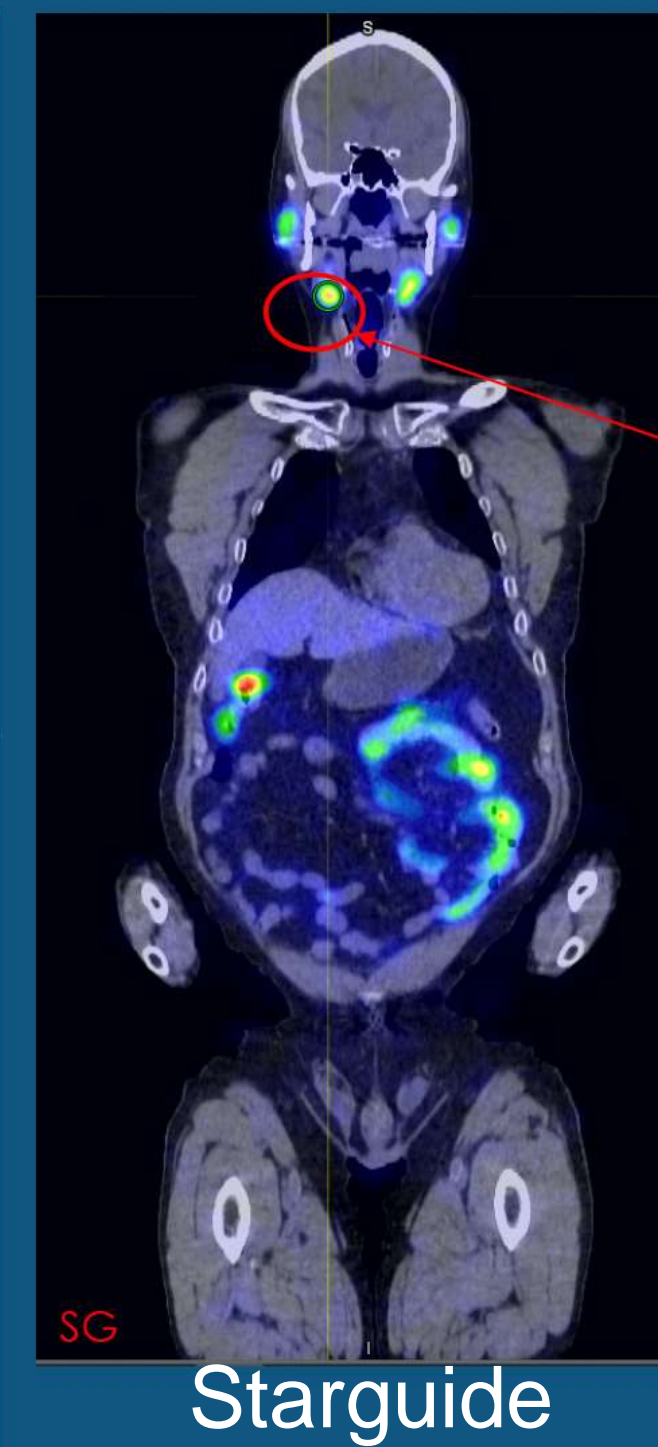
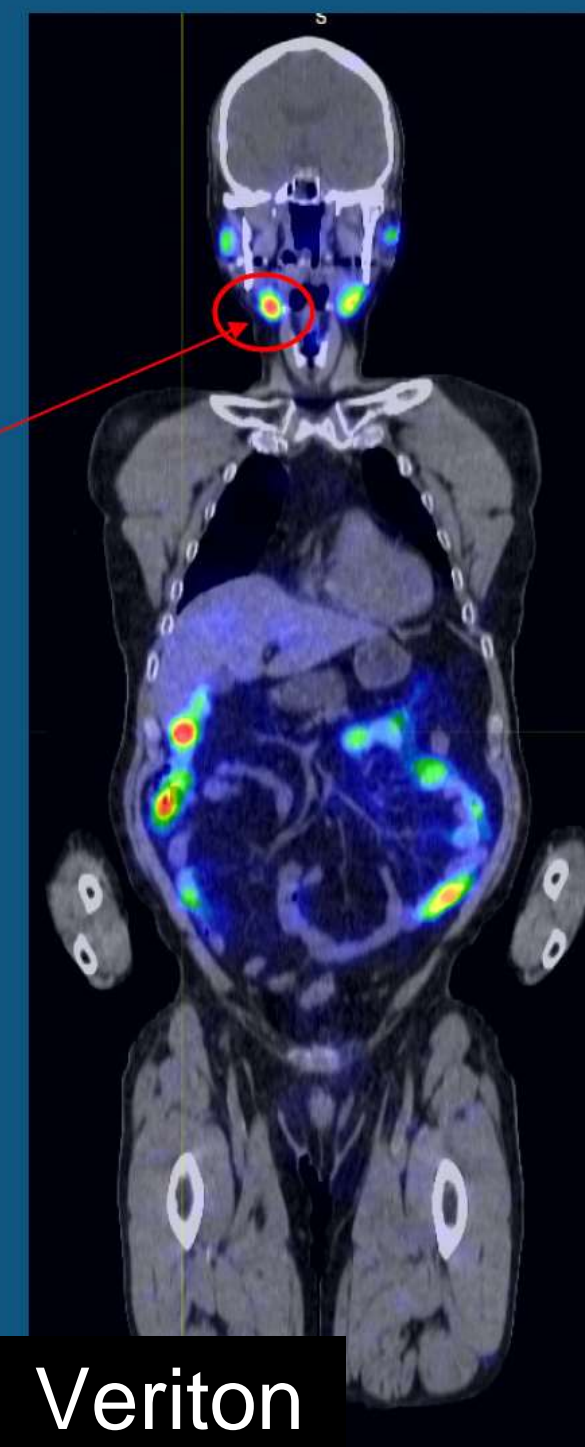


Images Courtesy of the Mayo Clinic, Rochester, MN, USA

Veriton or StarGuide? Pretty close

Veriton
 Max SUVbw = 4.4
 Mean SUVbw = 2
 Min SUVbw = 0.3

StarGuide
 Max SUVbw = 3.8
 Mean SUVbw = 1.7
 Min SUVbw = 0.4



Veriton
 Max SUVbw = 8.5
 Mean SUVbw = 5.2
 Min SUVbw = 3

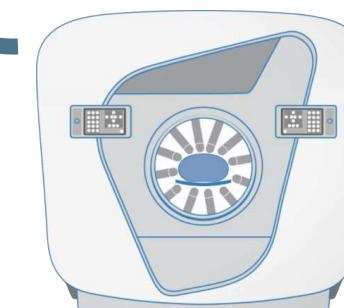
StarGuide
 Max SUVbw = 7.9
 Mean SUVbw = 5.5
 Min SUVbw = 3.4



CZT detector technology enables innovative SPECT system design

Full-Ring CZT SPECT/CT,

- ✓ Offers improved **sensitivity, energy resolution, and spatial resolution** compared to conventional SPECT systems leading to **enhanced imaging performance**
- ✓ Permits **quantitative rapid whole-body SPECT/3D imaging essential in high volume practice**
- ✓ Enables **3D dynamic imaging**
- ✓ Has the potential to deliver more **precise quantitative imaging** compared to conventional SPECT/CT systems with promising application in **theranostics for treatment monitoring and personalized dosimetry**
- ✓ Provides **adaptive data acquisition for improved imaging in a broad range of clinical applications**

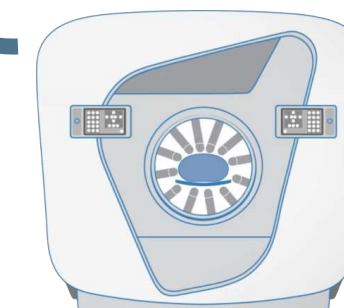




CZT detectors are becoming more common and less expensive to manufacture. However, still significantly more expensive than conventional SPECT/CT systems.

Established clinical applications, including *cardiac, liver, bone, theranostics, and brain*, now **justify the additional cost**. These **specialized applications** are taking **full-advantage of full-ring CZT system capabilities** (i.e., **Rapid Scan Time, Quantitation, and 3D Dynamic**). Exciting new possibilities for the future!

Conventional SPECT-Only systems will likely persist in the future. **Lower cost and sufficient for certain applications** (*e.g. gastric emptying, thyroid, and renal studies*)

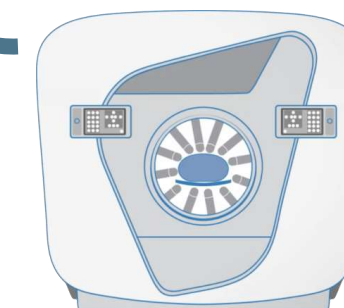




Absolute Quantitation Cross-Calibration is easier than with **conventional SPECT** (*fewer technical factors*). Uniform Flood Phantom leads to the best performance. **Jaszczak Phantom** provides a **good alternative** in routine practice for ^{99m}Tc .

However, these are not suited for **long-lived nuclides**. **Need for a robust alternative.**

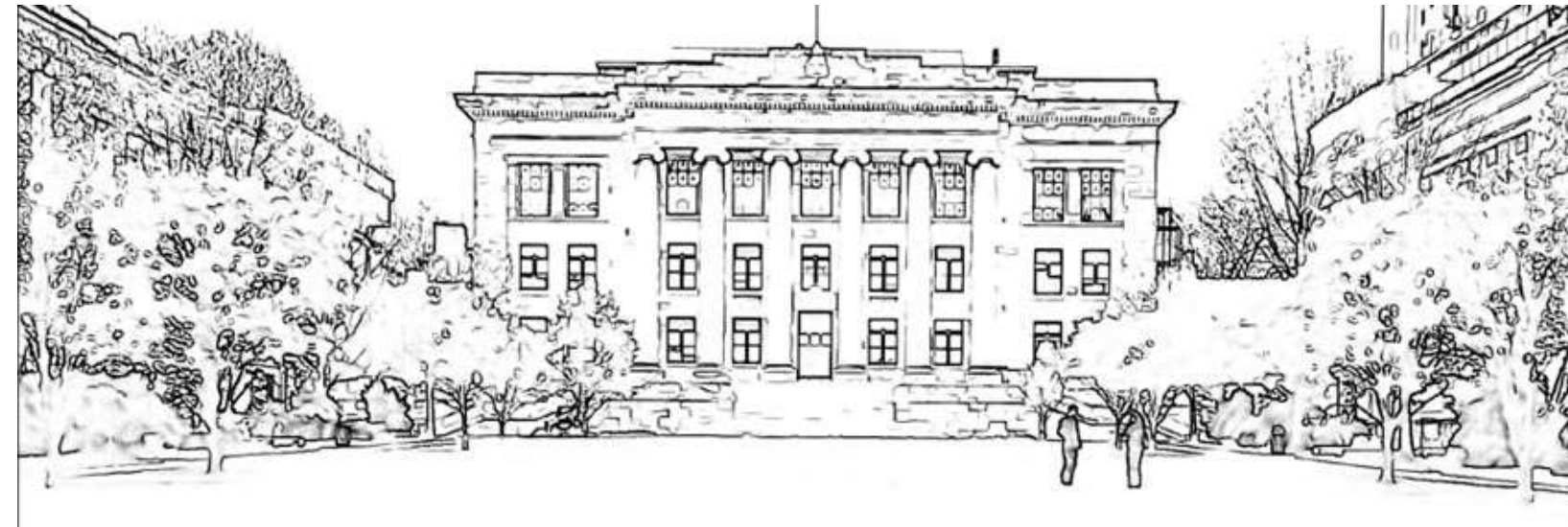
Need to assess **impact of parameters** on **absolute quantitation** and **establish guidelines**.





2023 CCSNMMI FALL EDUCATION PROGRAM
OCTOBER 21-22, 2023
NOBLESVILLE, IN

Latest Developments in CZT SPECT Imaging
B. Auer – April 14th, 2024



THANK YOU FOR YOUR ATTENTION

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