

Framework for Theranostic Digital Twins Generation and Virtual Theranostic Trials

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INTRODUCTION

Radiopharmaceutical therapy is established in metastatic castration-resistant prostate cancer. It is given as a fixed administered activity, not tailored to the absorbed dose reaching tumors and organs at risk.

Developing personalized planning needs data that is hard to obtain clinically:

- Serial quantitative SPECT with repeated scans over several days and absorbed dose map
- Prospective trials are costly and carry patient risk
- In real patients the absorbed dose is never truly known

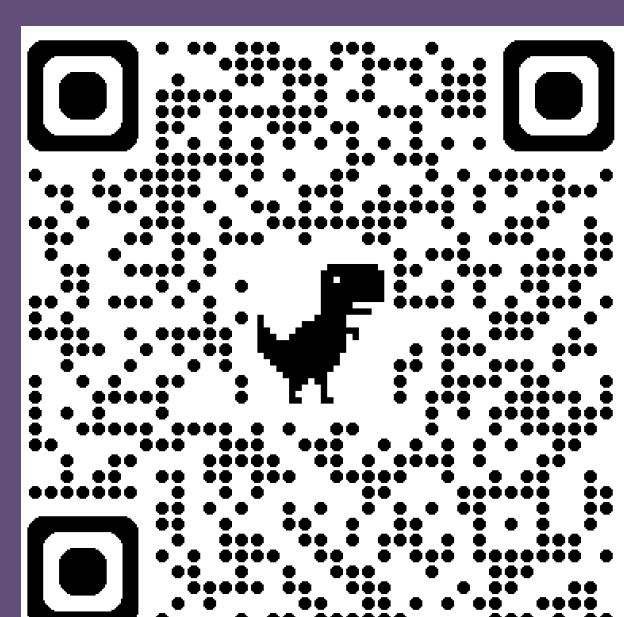
PyTheraTwin is an open-source tool that builds this dataset from patient anatomy alone. It generates synthetic SPECT images and absorbed dose maps from a prescribed activity distribution, so the input activity is known exactly. The framework produces complete synthetic datasets and supports virtual theranostic trials.

CONTACT

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<https://github.com/qurit/PyTheraTwin>



METHODS AND MATERIALS

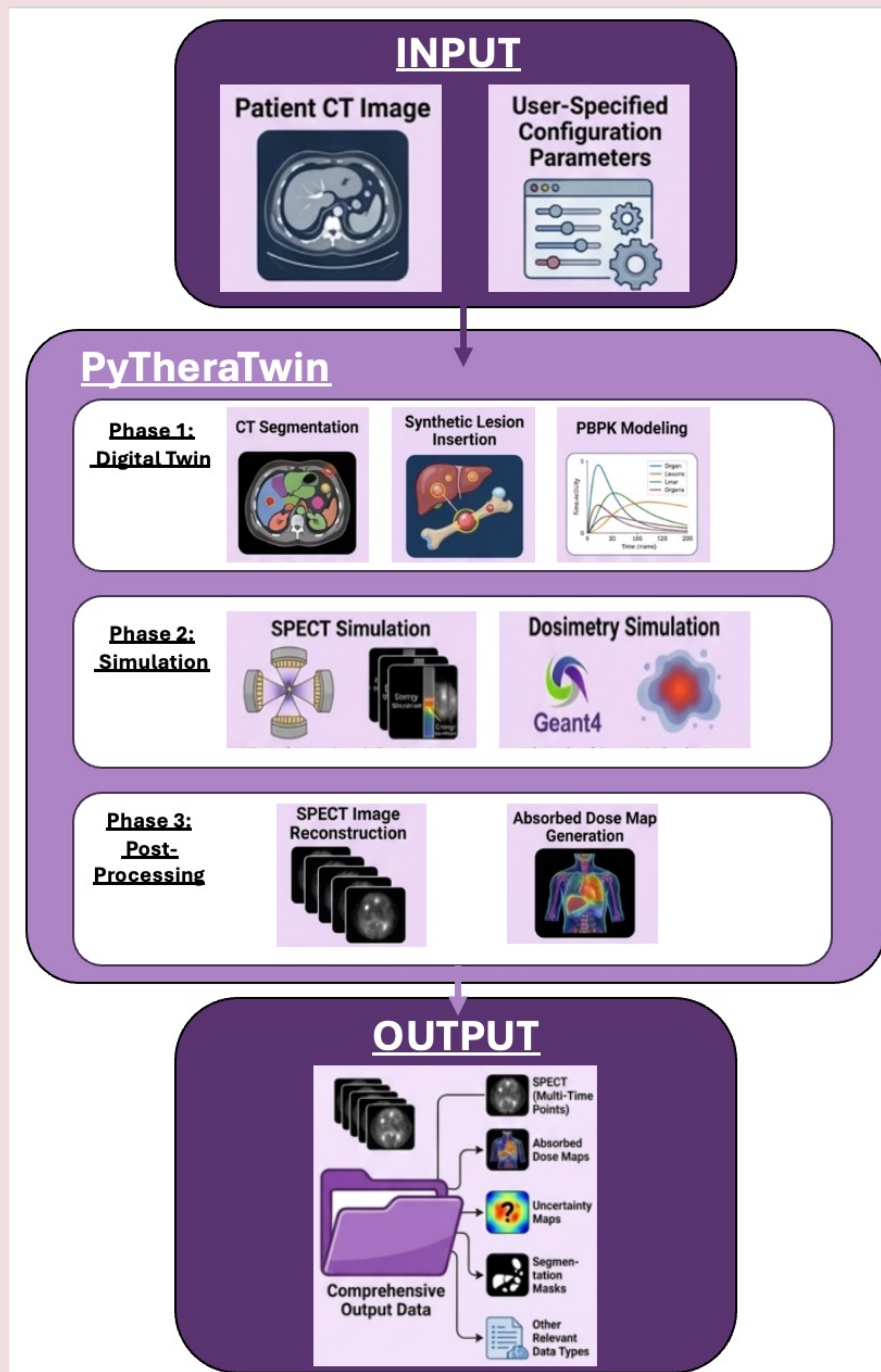


Figure 1. Framework workflow, from patient CT to synthetic SPECT images and absorbed dose maps.

	Parameter	Value
SPECT & Absorbed Dose	CPU Used	112 Cores
	XYZ Spacing [mm]	4.8, 4.8, 4.8
	Radioisotope	Lu-177
	Simulated ROIs	Kidney, Liver, Salivary Glands, Lungs, Synthetic Lesion(s), Remaining Body
SPECT Specific	# Photons	1e9
	# Projections	96
	Energy Window Width [%]	20 (+/- 10)
	OSEM (Iterations & Subset)	4 & 8
	Frame Times [hrs]	1, 2, 3, 8, 12, 24, 48, 72, 96, 120, 144, 168
Absorbed Dose Map Specific	Histories Per Batch	1e6
	Target Uncertainty [%]	5
	Electron Cut Off [mm]	0.5
Synthetic Lesions Parameters		
	Liver	Bone
Patient Number	11, 15, 17, 18	13, 16, 20
		Liver & Bone
		12, 14, 19

Table 1. Key framework configuration parameters for the 20-patient cohort.

RESULTS

DIGITAL TWIN

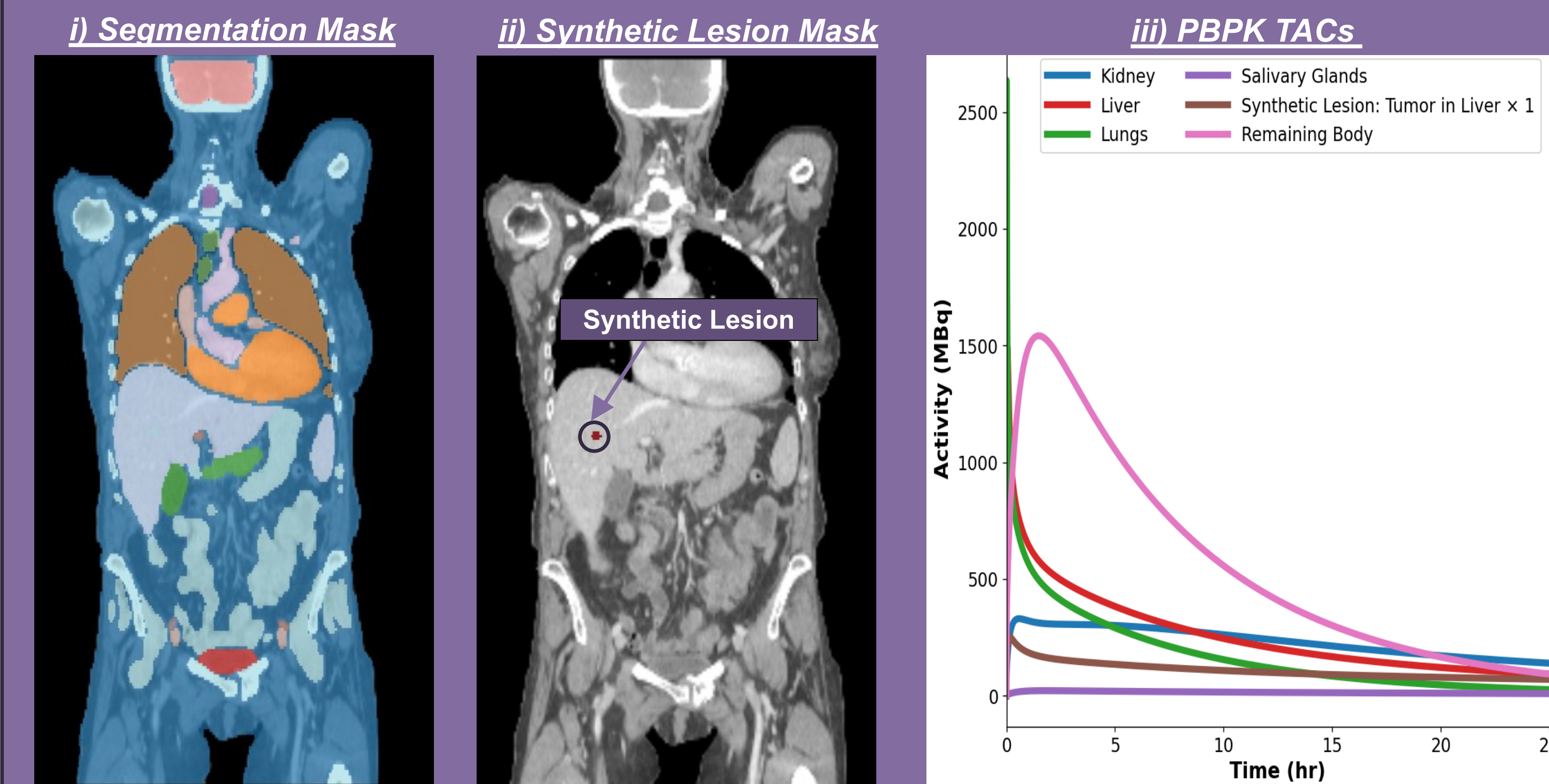


Figure 2. Phase 1 Outputs for Patient 11.

SIMULATION OUTPUTS

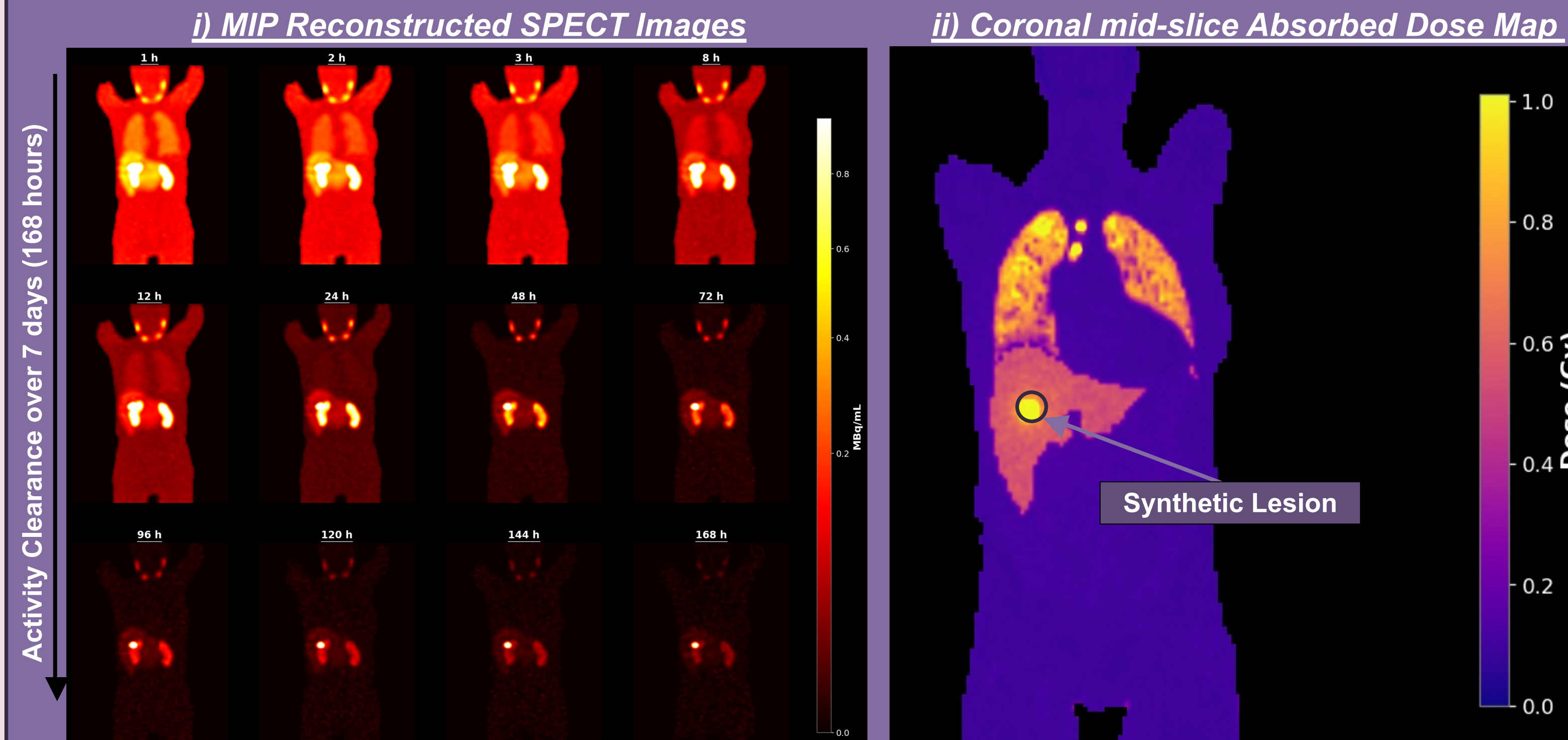


Figure 3. Phase 2 & 3 Outputs for Patient 11.

CONCLUSION

The framework builds patient-specific digital twins and simulates the synthetic SPECT images and absorbed dose maps that follow from a prescribed activity distribution. Because the input activity is modelled rather than measured, the simulated images and dose maps come with an exact reference that real patient scans never have.

We are releasing a complete 20-patient dataset of ground-truth absorbed dose, organ segmentations, and multi-time-point SPECT reconstructions with their activity distributions, which supports:

- Benchmarking dosimetry algorithms against realistic ground-truth absorbed dose maps
- Augmenting deep learning training for theranostics, such as lesion detection and tracking
- Generating realistic synthetic data for Virtual Theranostic Trials

Future work:

- Validate simulation realism against clinical ¹⁷⁷Lu-PSMA SPECT
- Add synthetic PET imaging and extend synthetic lesion realism
- Develop a predictive dosimetry model that is trained on PyTheraTwin outputs

REFERENCES

- [1] Wasserthal J, Breit HC, Meyer MT, et al. TotalSegmentator: robust segmentation of 104 anatomic structures in CT images. *Radiol Artif Intell*. 2023;5(5):e230024.
- [2] Felle-Parang A, Saboury B, Uribe C, Rahmim A. Physiologically based radiopharmacokinetic (PBPK) modeling to simulate and analyze radiopharmaceutical therapies. *EJNMMI Radiopharm Chem*. 2024;9:6.
- [3] Luekerberg M, Strand SE. A Monte Carlo program for the simulation of scintillation camera characteristics. *Comput Methods Programs Biomed*. 1989;25(4):257-272.
- [4] Samra D, Ashor N, Bauder T, et al. The OpenGATE ecosystem for Monte Carlo simulation in medical physics. *Phys Med Biol*. 2022;67(18):184001.
- [5] Polson LA, Fedirgo R, Li C, et al. PyTorchography: a Python library for medical image reconstruction. *SoftwareX*. 2025;29:102020.
- [6] Fedirgo R, Polson L, Li C, et al. Development of a theranostic digital twins framework to perform quantitative image analysis of radiopharmaceutical biodistributions. *J Nucl Med*. 2023;64(suppl 1).