

# Particle versus photon imaging for proton radiotherapy - an experimental comparison

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\* The first three authors contributed equally.

# Which imaging modality provides the highest RSP accuracy?

- Photon CT
  - 1) Single-energy CT
  - 2) Dual-energy CT
- Particle CT
  - 3) Proton CT
  - 4) Helium CT

# Which imaging modality provides the highest RSP accuracy?

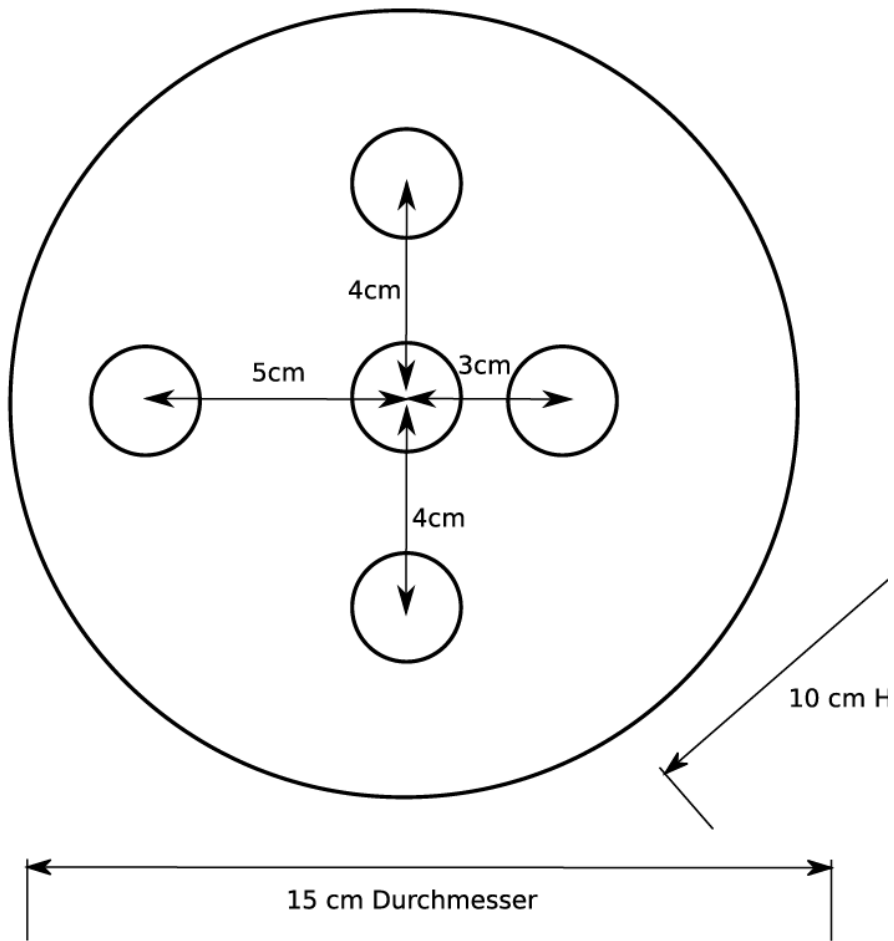
- 1) Design a **tissue phantom**, collect and prepare tissue samples.
- 2) Perform **reference RSP** measurements.
- 3) Photon CT imaging: Collect **SECT and DECT** images, estimate the RSPs and compare to reference RSP.
- 4) Particle CT imaging: Collect **proton and helium CT** images, compare RSPs to photon CT and reference RSP.

All in one  
day!!

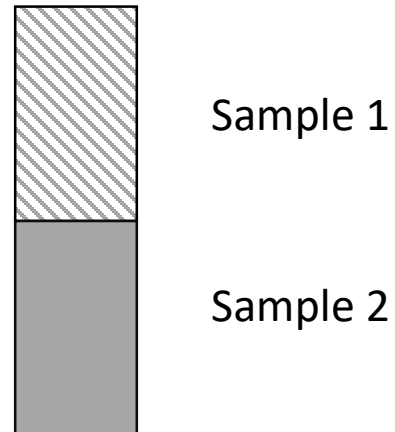
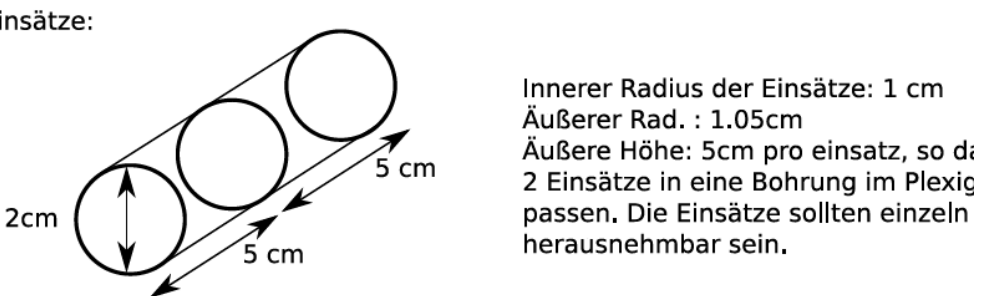
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# Tissue phantom design



- PMMA cylinder
  - 15 cm diameter
  - 10 cm high
  - Cylindrical holes to insert 3D printed containers
  - Holds 10 containers in one scan





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# Tissue sample preparation

- 17 porcine and bovine samples including:
  - Lung
  - Fat
  - Marrow
  - Blood
  - Muscle
  - Brain
  - Kidney
  - Liver
  - Trabecular bone
  - Cortical bone







# Tissue sample preparation

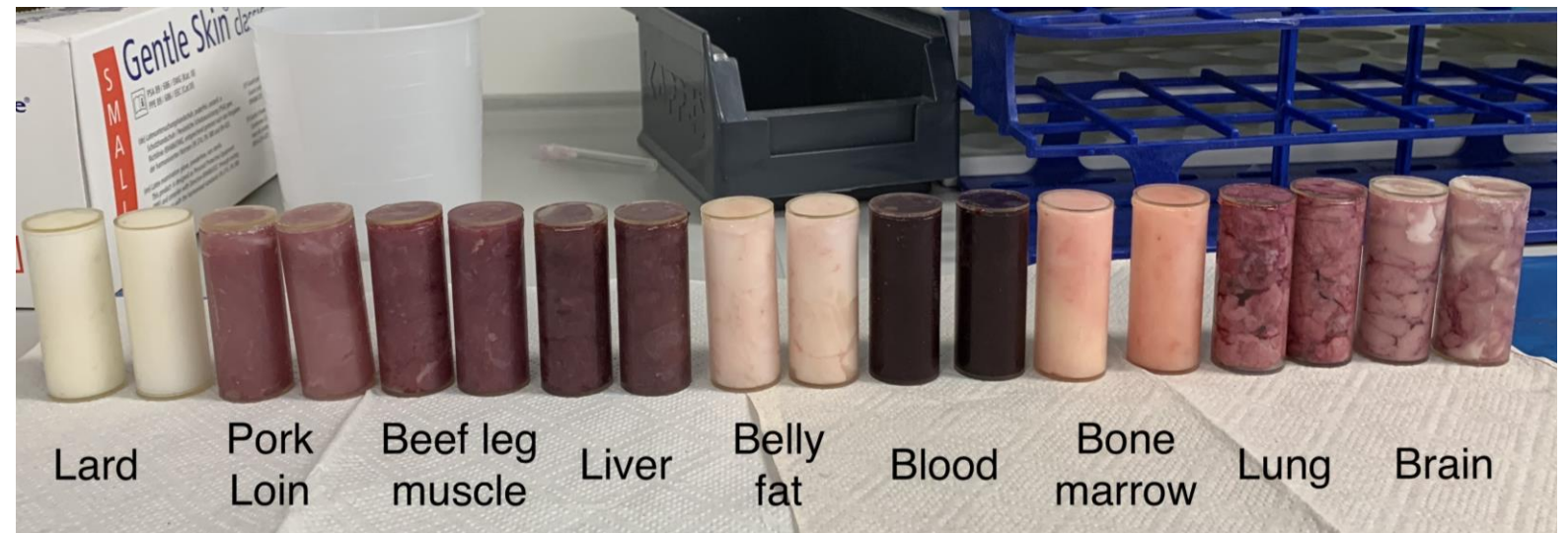
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# Tissue sample preparation

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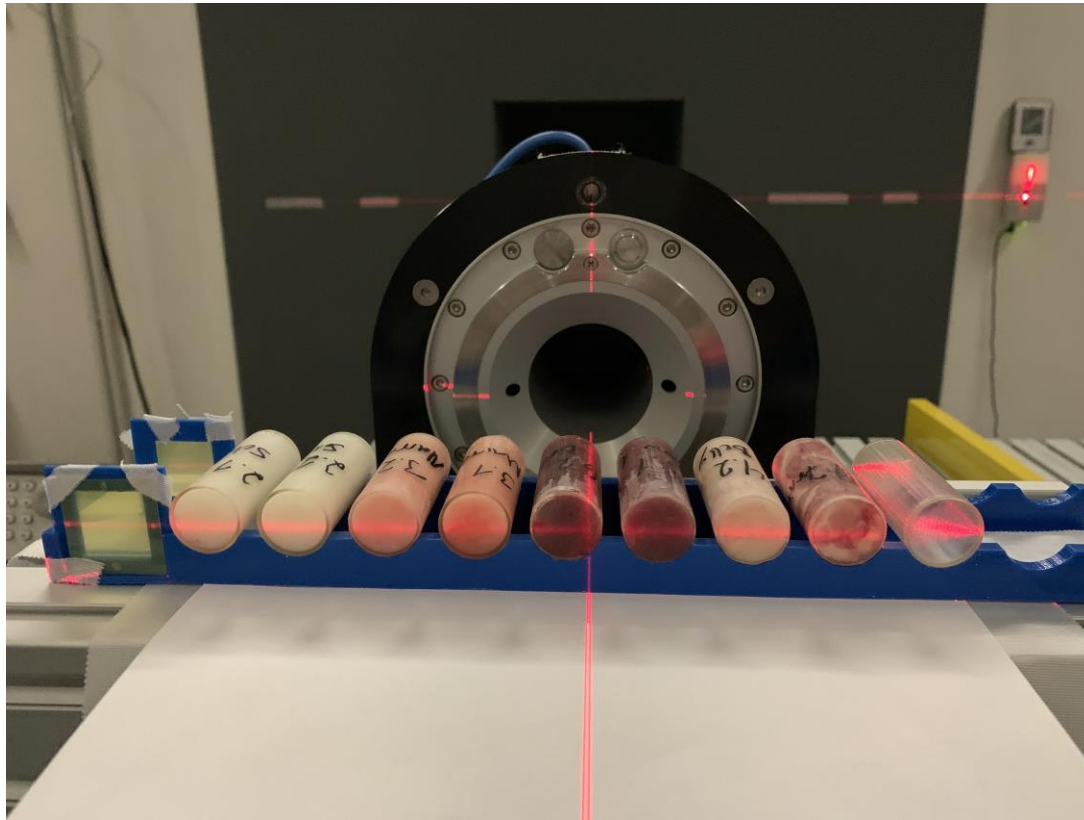
- Lung
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# Which imaging modality provides the highest RSP accuracy?

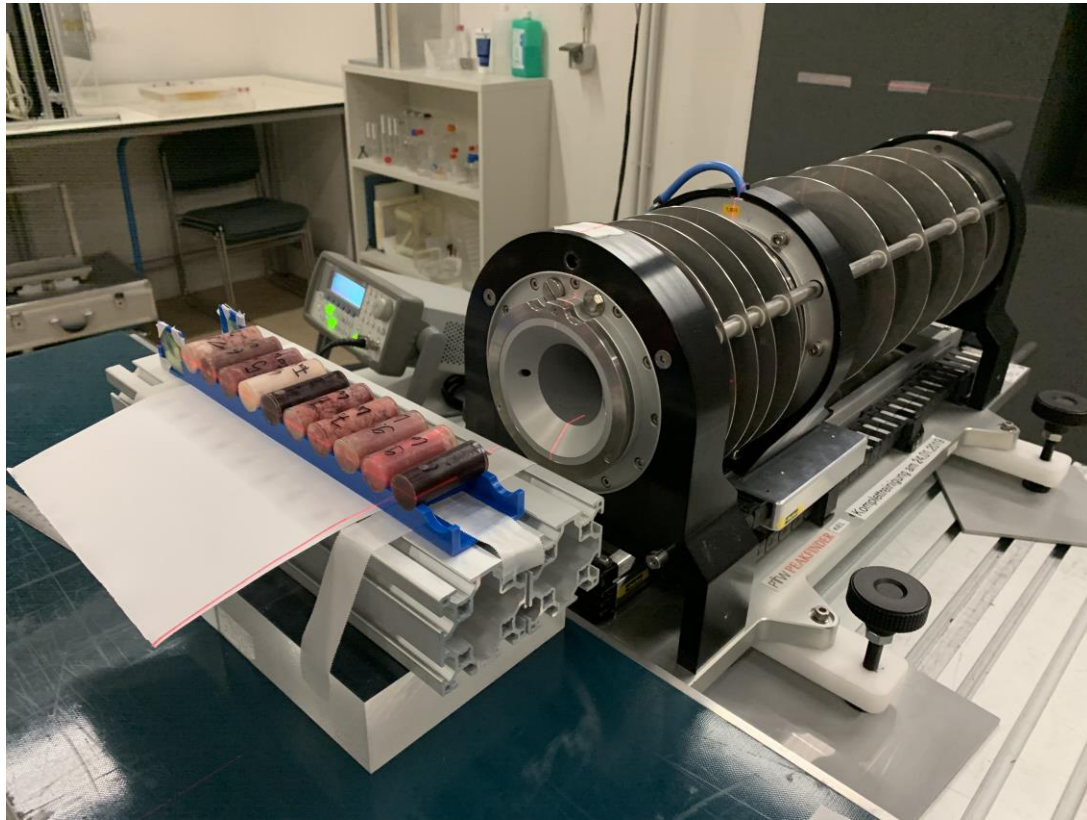
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# Reference RSP measurements



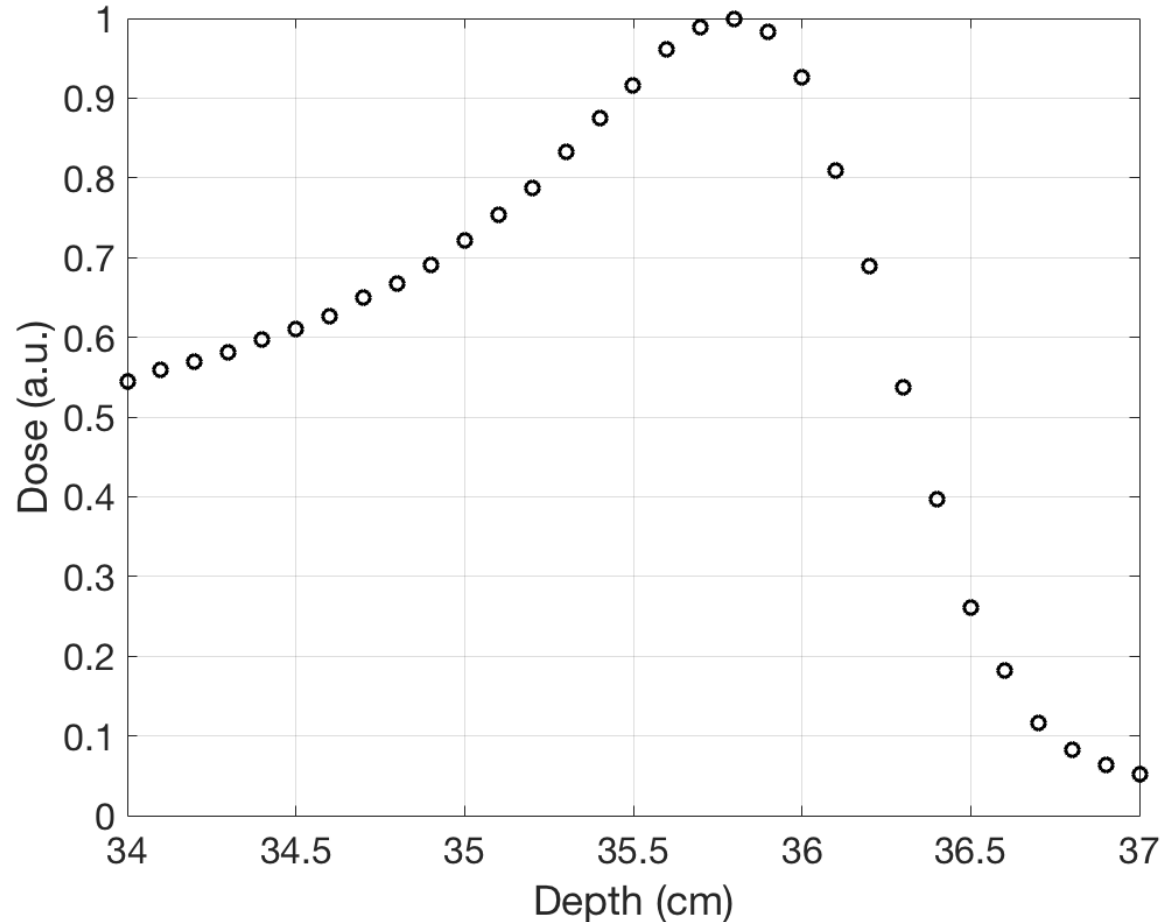
- Samples in 3D printed containers
  - Placed on a 3D printed holder and a translational stage

# Reference RSP measurements



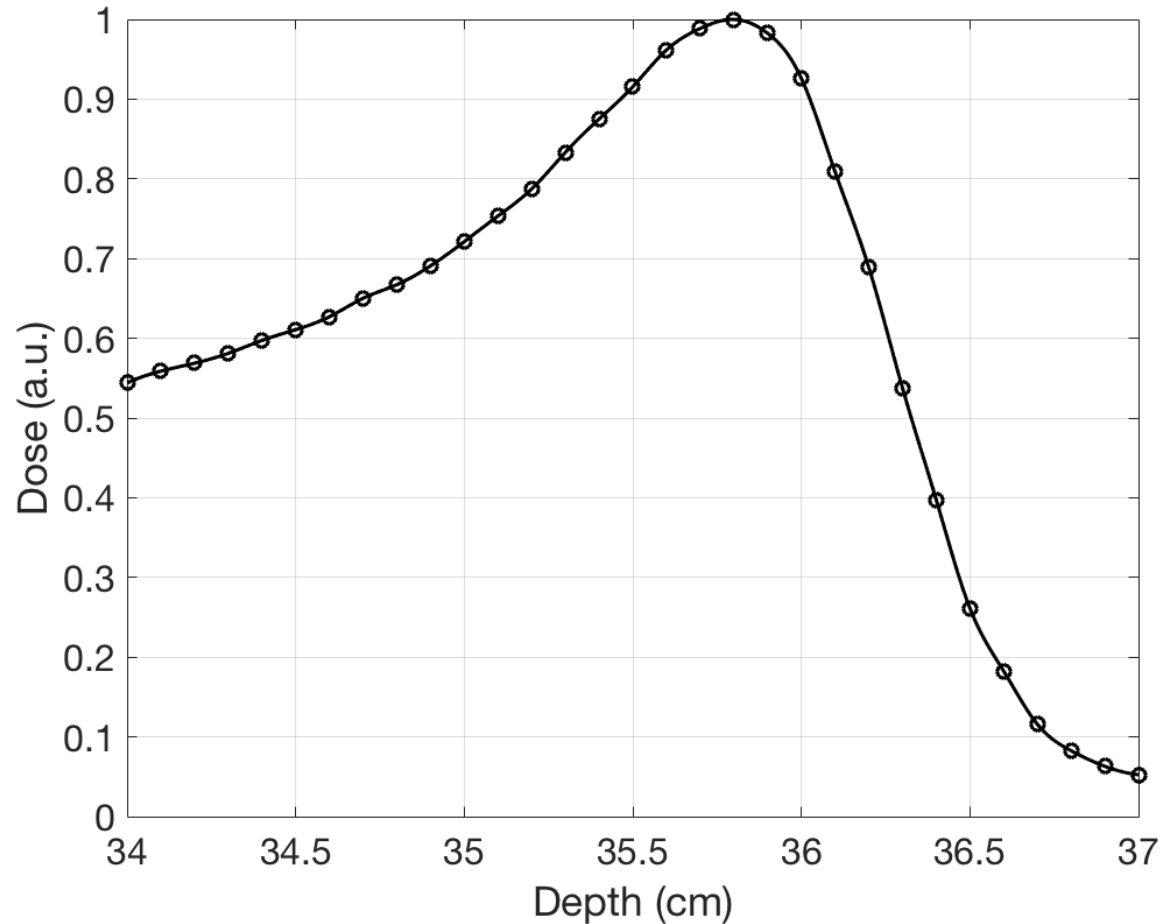
- Samples in 3D printed containers
  - Placed on a 3D printed holder and a translational stage
- PTW *PeakFinder*
  - Adjustable water column for peak detection measurements
- Clinical carbon beam
  - Narrow peak, low scattering

# Reference RSP measurements



- For each tissue sample and empty container: measure a depth dose curve

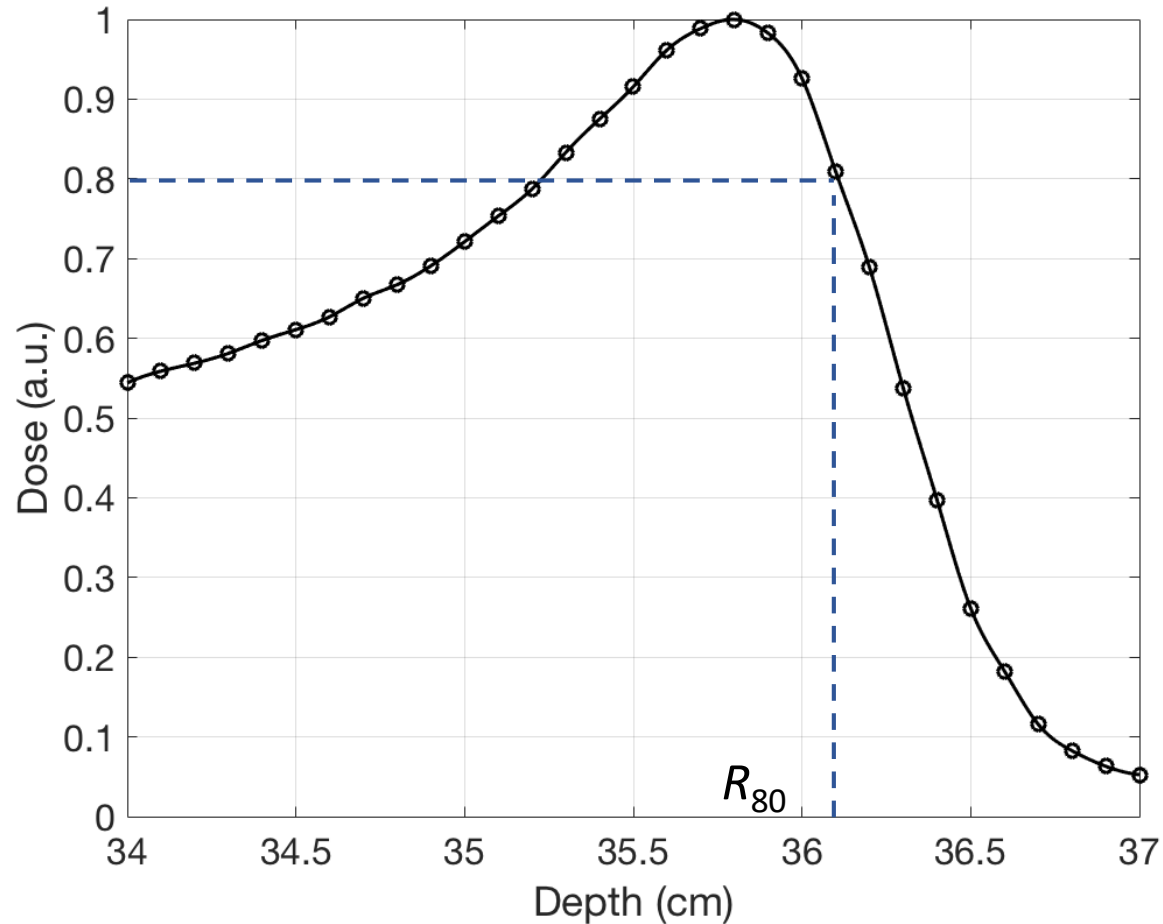
# Reference RSP measurements



- For each tissue sample and empty container: measure a depth dose curve
- Interpolate to find the 80% distal fall-off

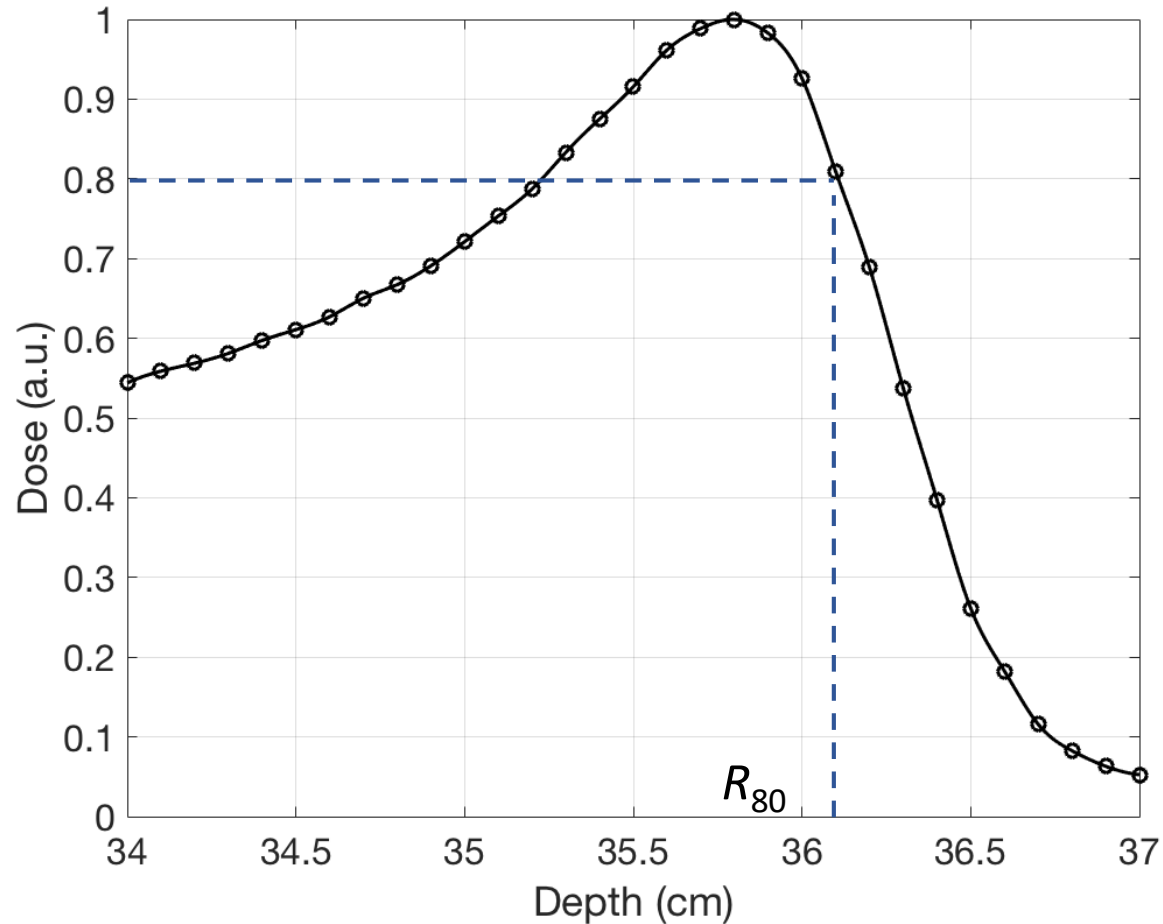


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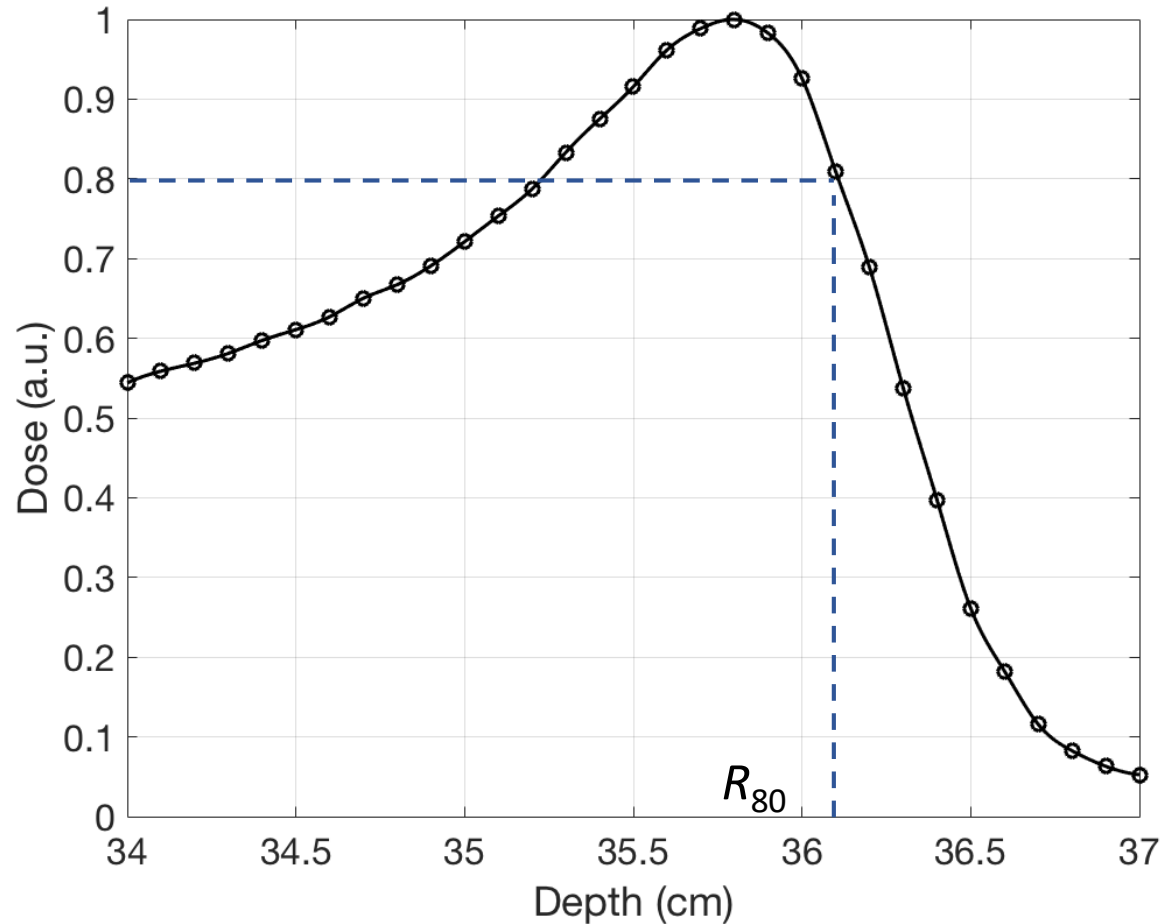
# Reference RSP measurements



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$$\text{WET}_{\text{sample}} = R_{80,\text{empty}} - R_{80,\text{sample}}$$

# Reference RSP measurements



- For each tissue sample and empty container: measure a depth dose curve
- Interpolate to find the 80% distal fall-off

$$\text{WET}_{\text{sample}} = R_{80,\text{empty}} - R_{80,\text{sample}}$$

$$\text{RSP}_{\text{sample,ref}} = \frac{\text{WET}_{\text{sample}}}{t}$$

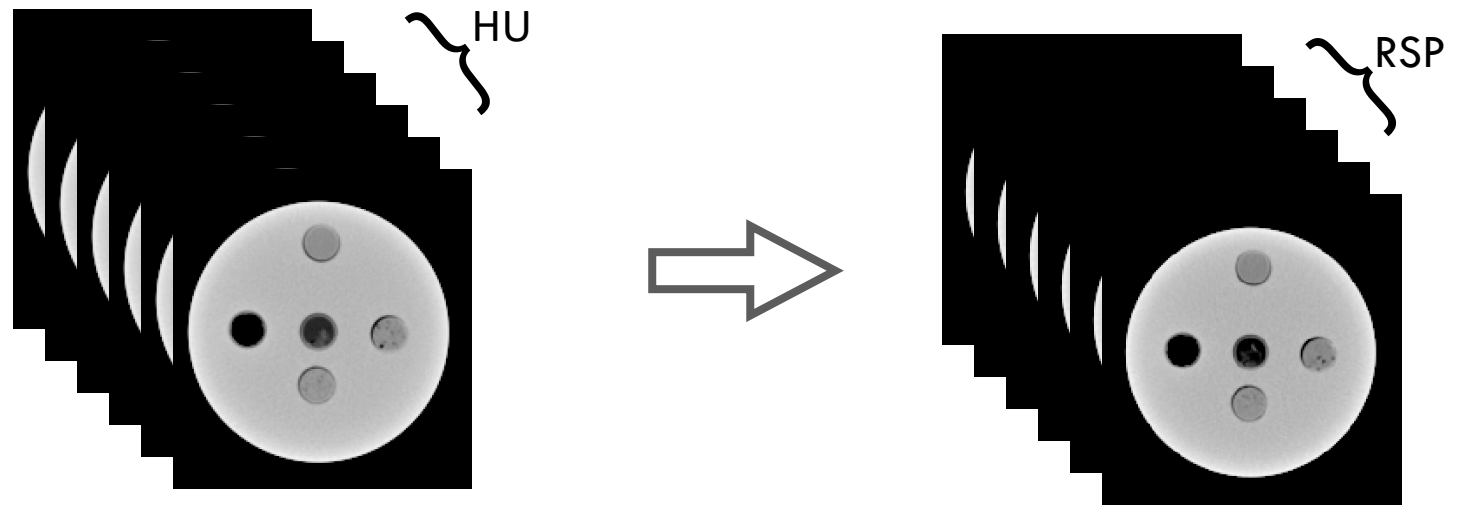
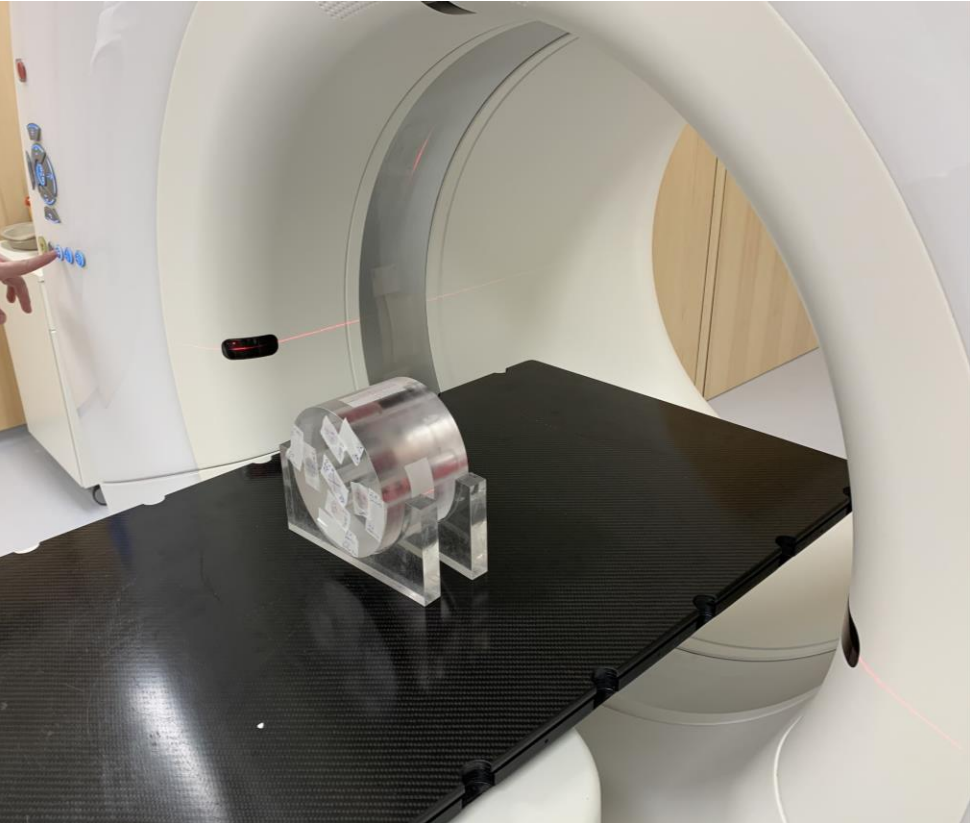
# Reference RSP measurements

Tissue	RSP <sub>ref</sub>
Lung	0.90 ± 0.04
Belly fat	1.00 ± 0.00
Back fat	0.97 ± 0.01
Marrow	0.93 ± 0.02
Water	1.00 ± 0.00
Blood	1.05 ± 0.00
Cheek muscle	1.05 ± 0.01
Loin 1	1.06 ± 0.00
Loin 2	1.06 ± 0.00
Leg muscle	1.05 ± 0.01
Brain	1.04 ± 0.00
Kidney 1	1.05 ± 0.00
Kidney 2	1.04 ± 0.01
Liver 1	1.06 ± 0.00
Liver 2	1.06 ± 0.00
Trabecular bone	1.19 ± 0.06
Cortical bone	1.78 ± 0.03
Mean error	–
Mean absolute error	–
Root mean square error	–

# Which imaging modality provides the highest RSP accuracy?

- 1) Design a **tissue phantom**, collect and prepare tissue samples.
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# Estimating RSP maps from SECT Images

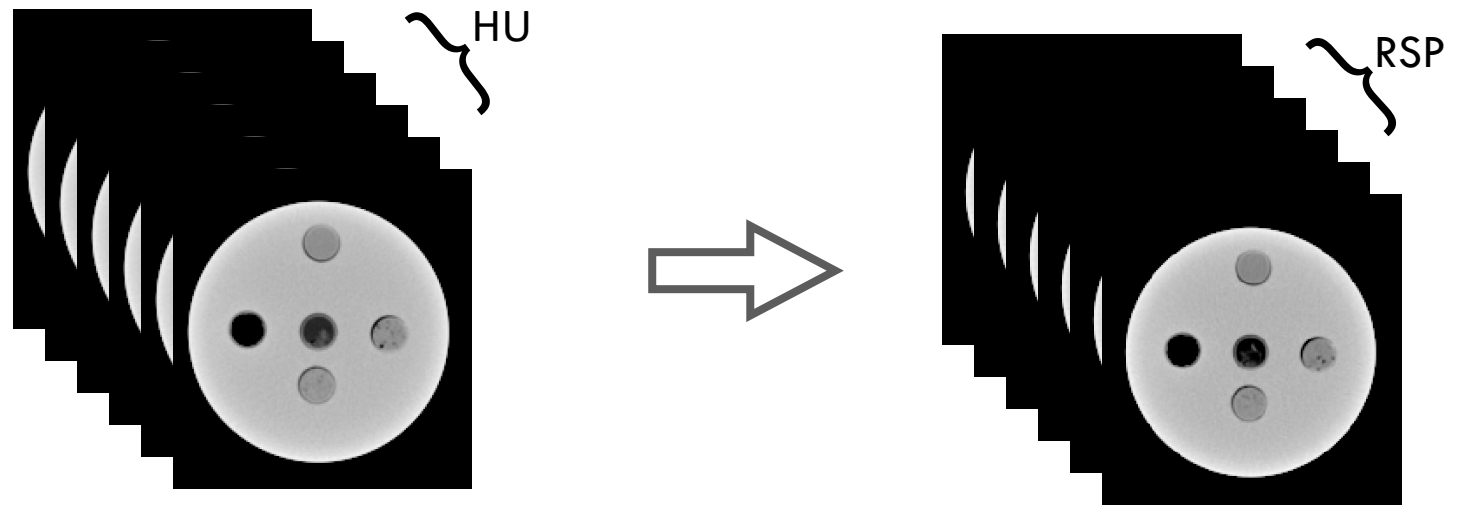
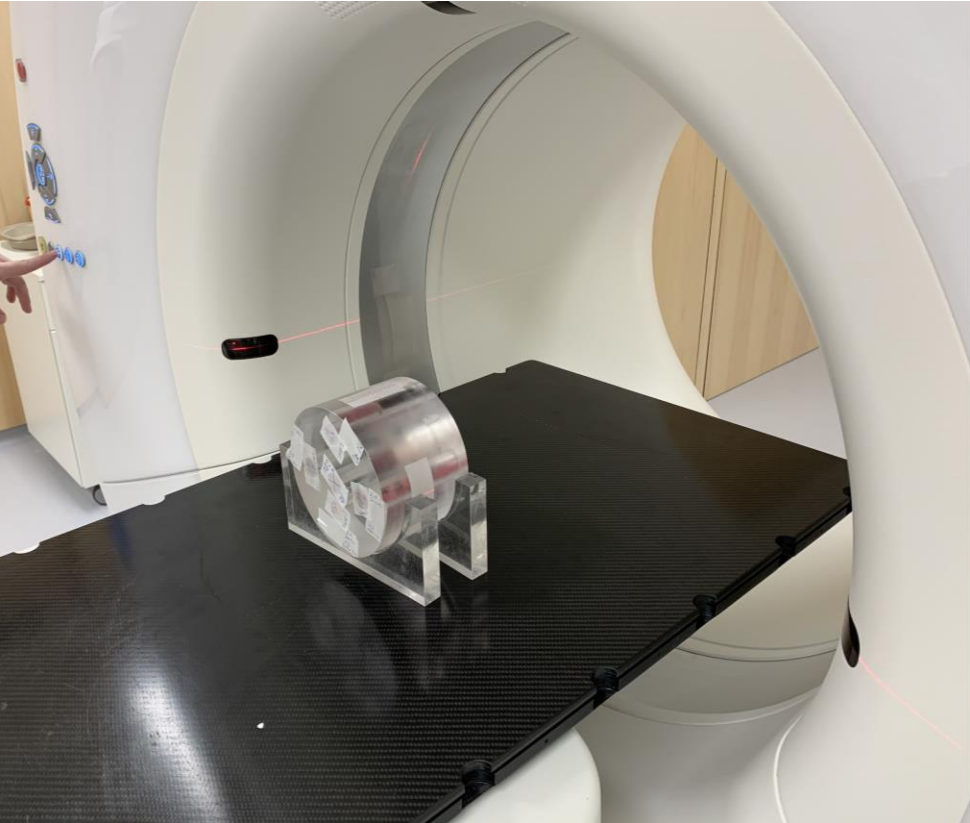


## Calibration

- Schneider, U., Pedroni, E. and Lomax, A., 1996. The calibration of CT Hounsfield units for radiotherapy treatment planning. *Physics in Medicine & Biology*, 41(1), p.111.
- Calibration phantom: Gammex RMI 467 electron density phantom
- Reference tissues: White and Woodard



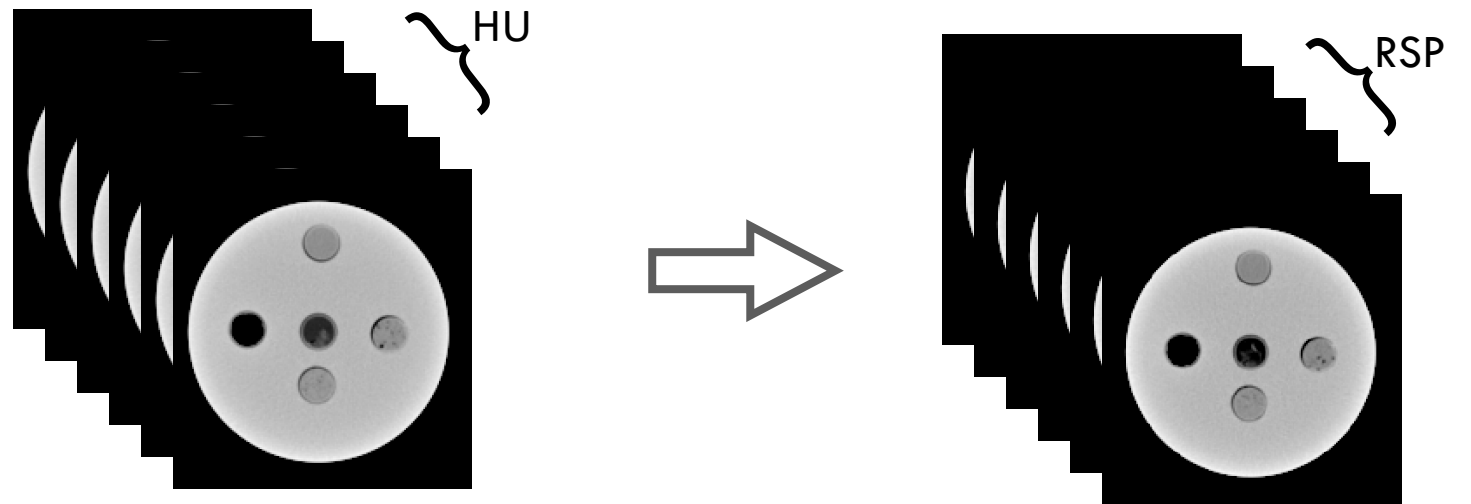
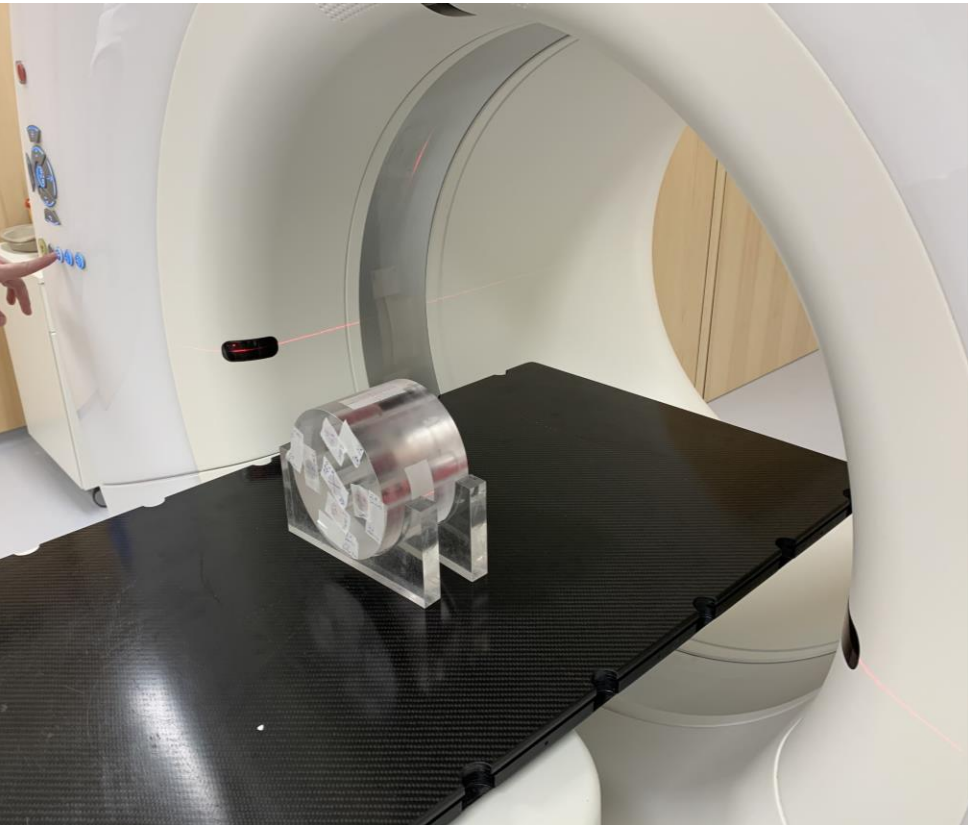
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- Measure average RSP per sample in VOIs:  $\mathbf{RSP}_{\text{SECT}}$

# Estimating RSP maps from SECT Images

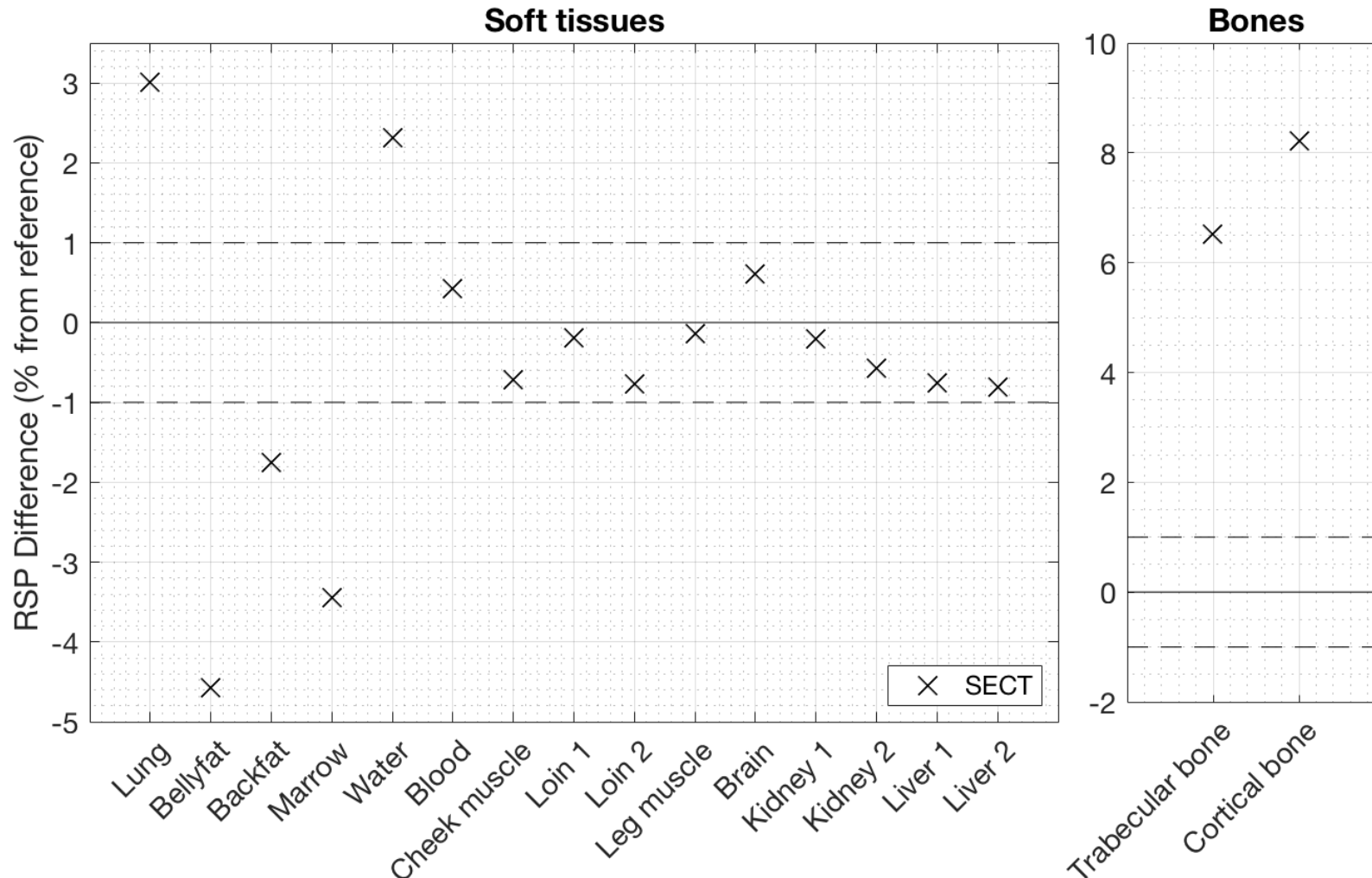


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- Calibration phantom: Gammex RMI 467 electron density phantom
- Reference tissues: White and Woodard
- Measure average RSP per sample in VOIs:  $\mathbf{RSP}_{\text{SECT}}$

$$\Delta\text{RSP}(\%) = \frac{\text{RSP}_{\text{SECT}} - \text{RSP}_{\text{ref}}}{\text{RSP}_{\text{ref}}} * 100$$

# SECT-estimated RSP errors

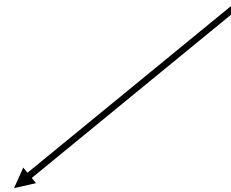


$$\Delta RSP(\%) = \frac{RSP_{SECT} - RSP_{ref}}{RSP_{ref}} * 100$$

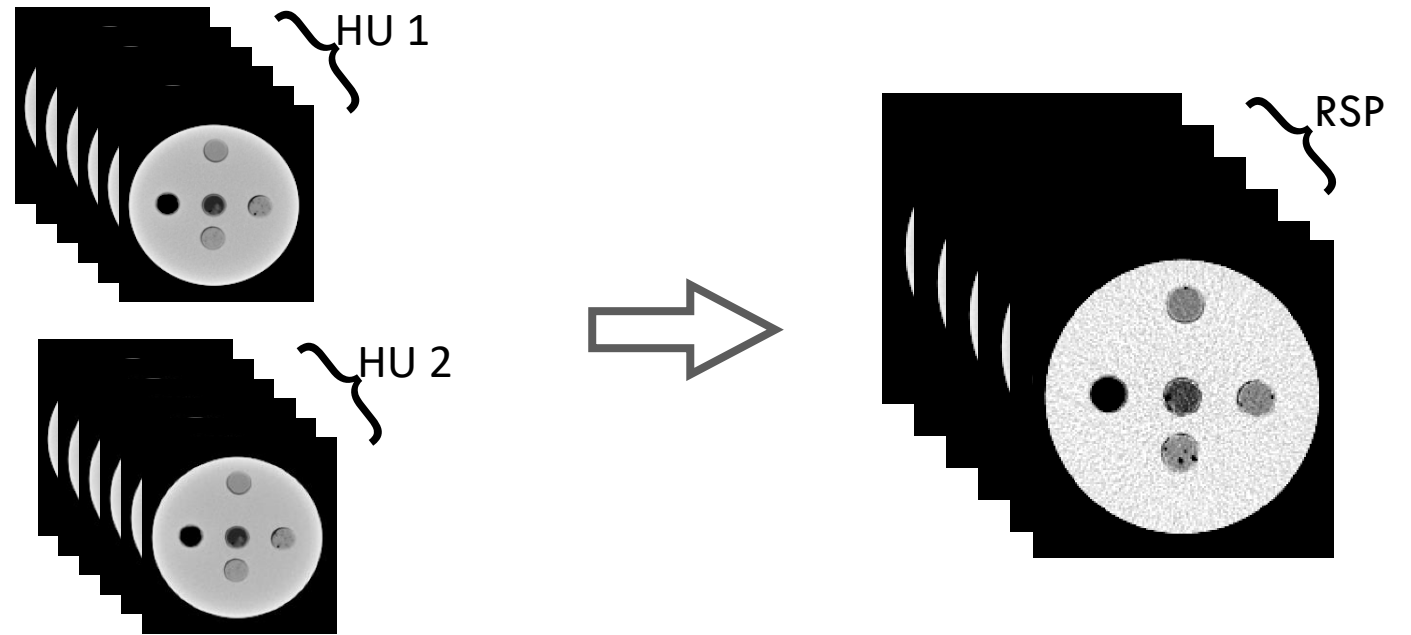
# SECT-estimated RSP errors

Tissue	RSP <sub>ref</sub>	SECT
Lung	0.90 ± 0.04	3.01 ± 11.06
Belly fat	1.00 ± 0.00	-4.57 ± 3.35*
Back fat	0.97 ± 0.01	-1.75 ± 1.97
Marrow	0.93 ± 0.02	-3.44 ± 5.78
Water	1.00 ± 0.00	2.32 ± 0.31
Blood	1.05 ± 0.00	0.43 ± 0.29
Cheek muscle	1.05 ± 0.01	-0.72 ± 1.49
Loin 1	1.06 ± 0.00	-0.20 ± 2.18
Loin 2	1.06 ± 0.00	-0.77 ± 1.51
Leg muscle	1.05 ± 0.01	-0.14 ± 1.68
Brain	1.04 ± 0.00	0.61 ± 0.83
Kidney 1	1.05 ± 0.00	-0.20 ± 3.25
Kidney 2	1.04 ± 0.01	-0.58 ± 4.90
Liver 1	1.06 ± 0.00	-0.75 ± 4.61
Liver 2	1.06 ± 0.00	-0.80 ± 6.31
Trabecular bone	1.19 ± 0.06	6.52 ± 6.86
Cortical bone	1.78 ± 0.03	8.22 ± 5.06 <sup>†</sup>
Mean error	–	<b>0.42 ± 4.55</b>
Mean absolute error	–	<b>2.06</b>
Root mean square error	–	<b>3.10</b>

$$\Delta\text{RSP}(\%) = \frac{\text{RSP}_{\text{SECT}} - \text{RSP}_{\text{ref}}}{\text{RSP}_{\text{ref}}} * 100$$



# Estimating RSP maps from DECT Images

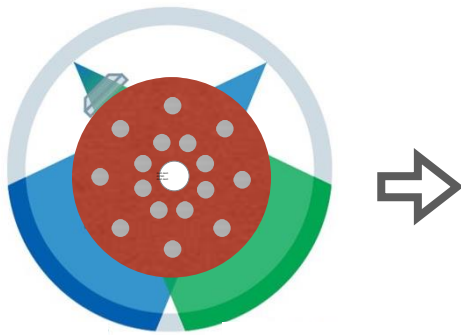


## Calibration

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# Estimating RSP maps from DECT Images

- Parameterize the **CT number**  $u$  as a function of  $Z_{\text{med}}$  and the electron density  $\rho_e$

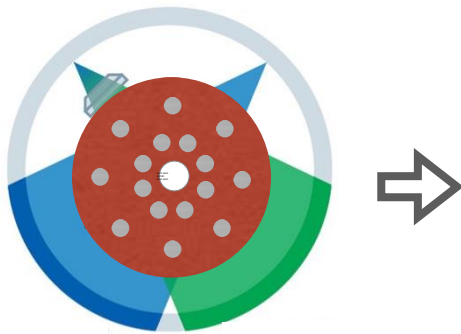


$$u = \rho_e \sum_{m=1}^M b_m Z_{\text{med}}^{m-1}$$



# Estimating RSP maps from DECT Images

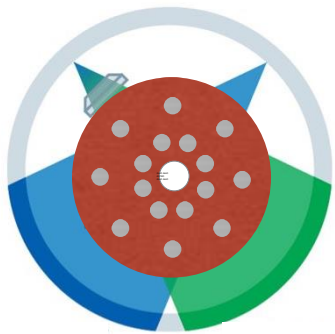
- Parameterize the **CT number**  $u$  as a function of  $Z_{\text{med}}$  and the electron density  $\rho_e$
- Parameterize the **effective atomic number**  $Z_{\text{med}}$  as a function of the dual energy index  $\Gamma$



$$u = \rho_e \sum_{m=1}^M b_m Z_{\text{med}}^{m-1}$$
$$Z_{\text{med}} = \sum_{k=1}^K c_k \Gamma^{k-1}$$

# Estimating RSP maps from DECT Images

- Parameterize the **CT number**  $u$  as a function of  $Z_{\text{med}}$  and the electron density  $\rho_e$
- Parameterize the **effective atomic number**  $Z_{\text{med}}$  as a function of the dual energy index  $\Gamma$
- **Calibrate** (find parameters  $c_k$  and  $b_m$ ) using tissue substitutes with known densities and compositions



$$u = \rho_e \sum_{m=1}^M b_m Z_{\text{med}}^{m-1}$$

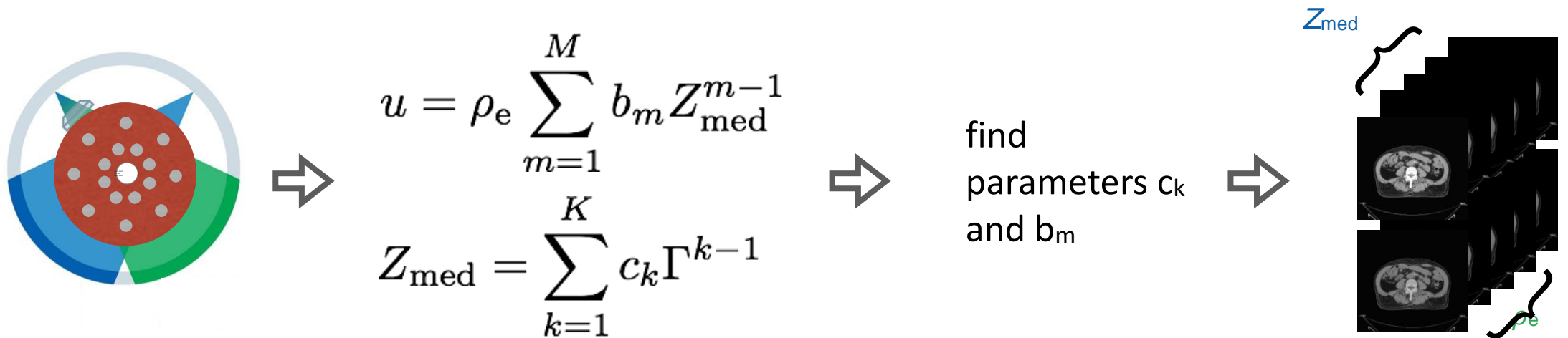
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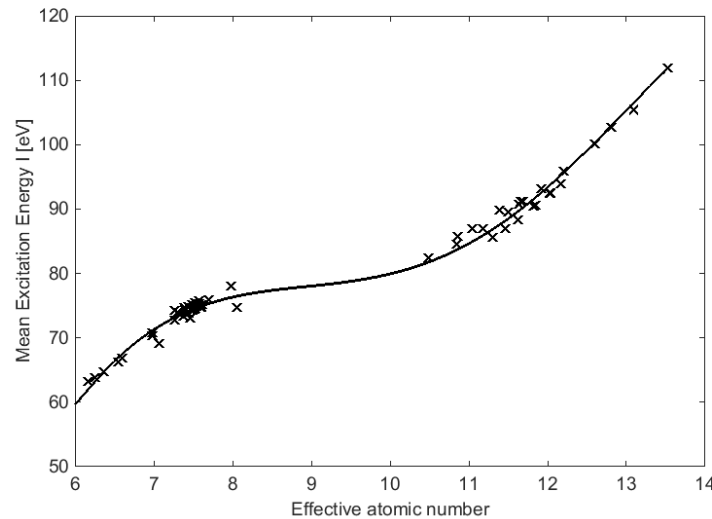
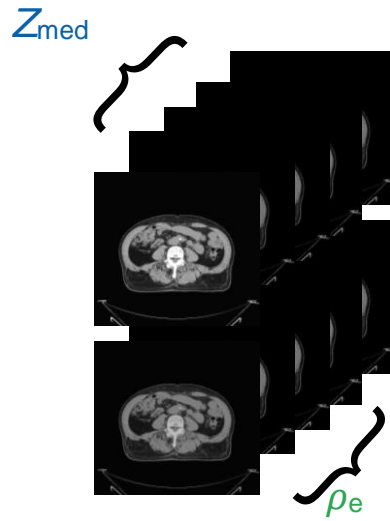
# Estimating RSP maps from DECT Images

- Parameterize the **CT number**  $u$  as a function of  $Z_{\text{med}}$  and the electron density  $\rho_e$
- Parameterize the **effective atomic number**  $Z_{\text{med}}$  as a function of the dual energy index  $\Gamma$
- **Calibrate** (find parameters  $c_k$  and  $b_m$ ) using tissue substitutes with known densities and compositions
- use parameters to find  $Z_{\text{med}}$  and  $\rho_e$  per voxel in a patient scan



# Estimating RSP maps from DECT Images

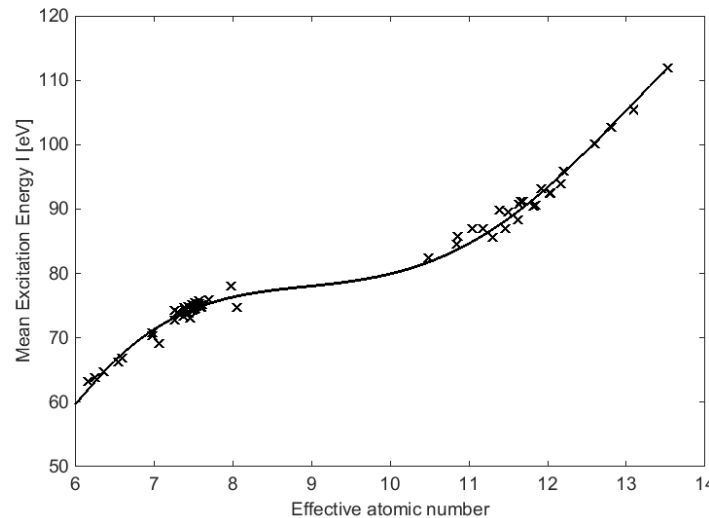
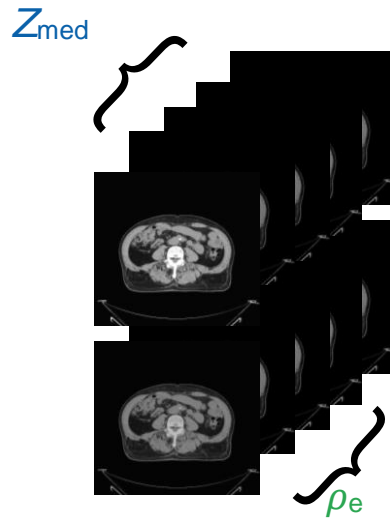
- Parameterize the **mean excitation energy  $I$**  as a function of  $Z_{\text{med}}$



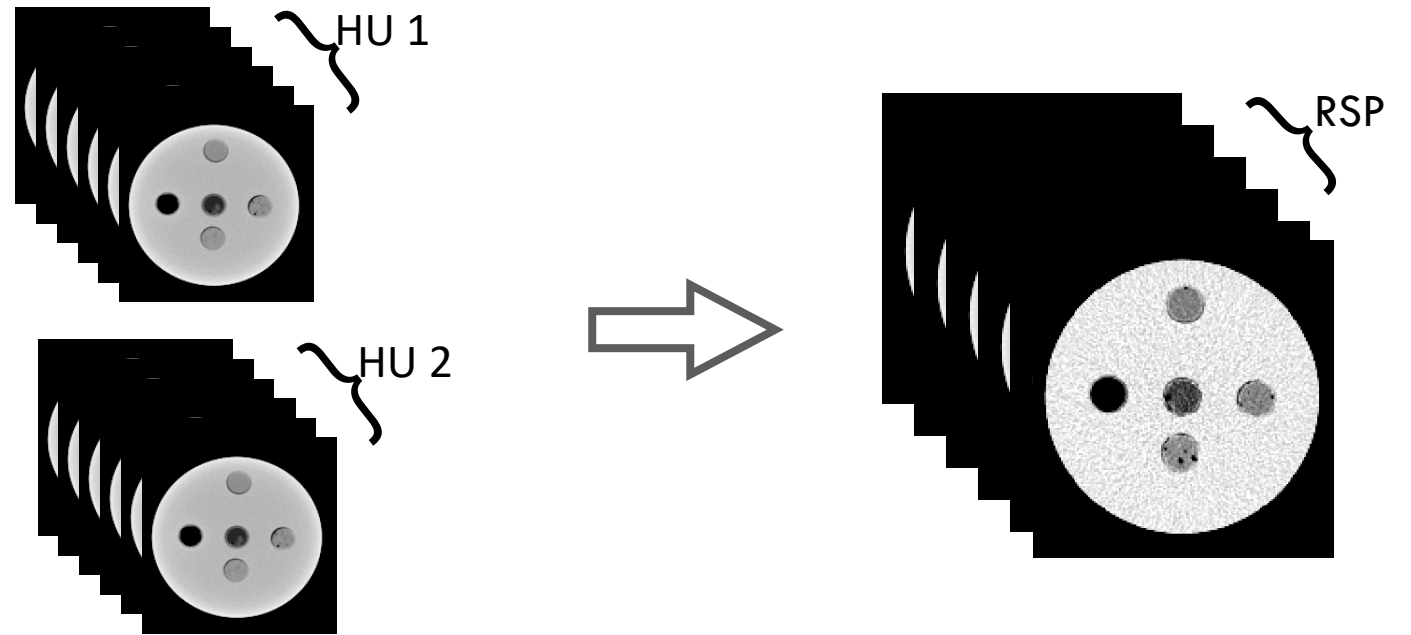
# Estimating RSP maps from DECT Images

- Parameterize the **mean excitation energy  $I$**  as a function of  $Z_{\text{med}}$
- Calculate the **RSP** from  $I$  and  $\rho_e$  per voxel using the **Bethe equation**

$$-\frac{dE}{dx} = \rho_e \frac{k_0 z^2}{\beta^2} \left[ \ln \left( \frac{2m_e c^2 \beta^2}{I(1 - \beta^2)} \right) - \beta^2 \right]$$



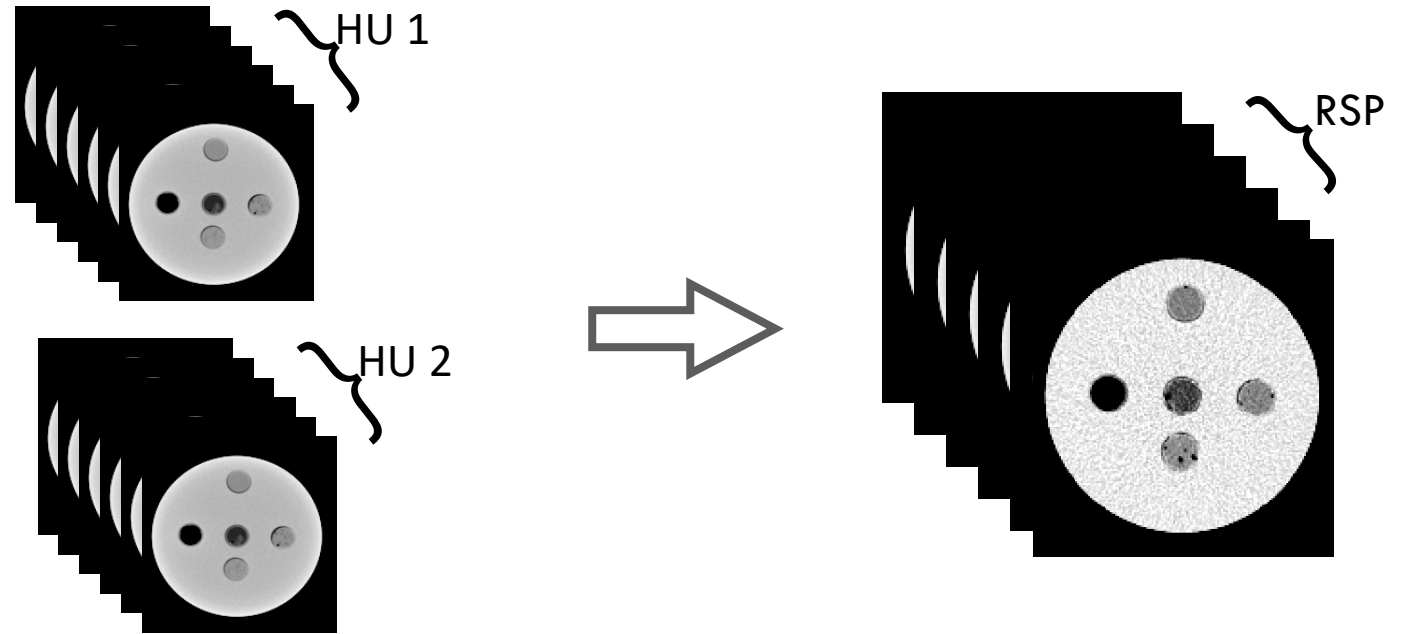
# Estimating RSP maps from DECT Images



## Calibration

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- Calibration phantom: Gammex RMI 467 electron density phantom
- Measure average RSP per sample in VOIs:  $\mathbf{RSP}_{\text{DECT}}$

# Estimating RSP maps from DECT Images



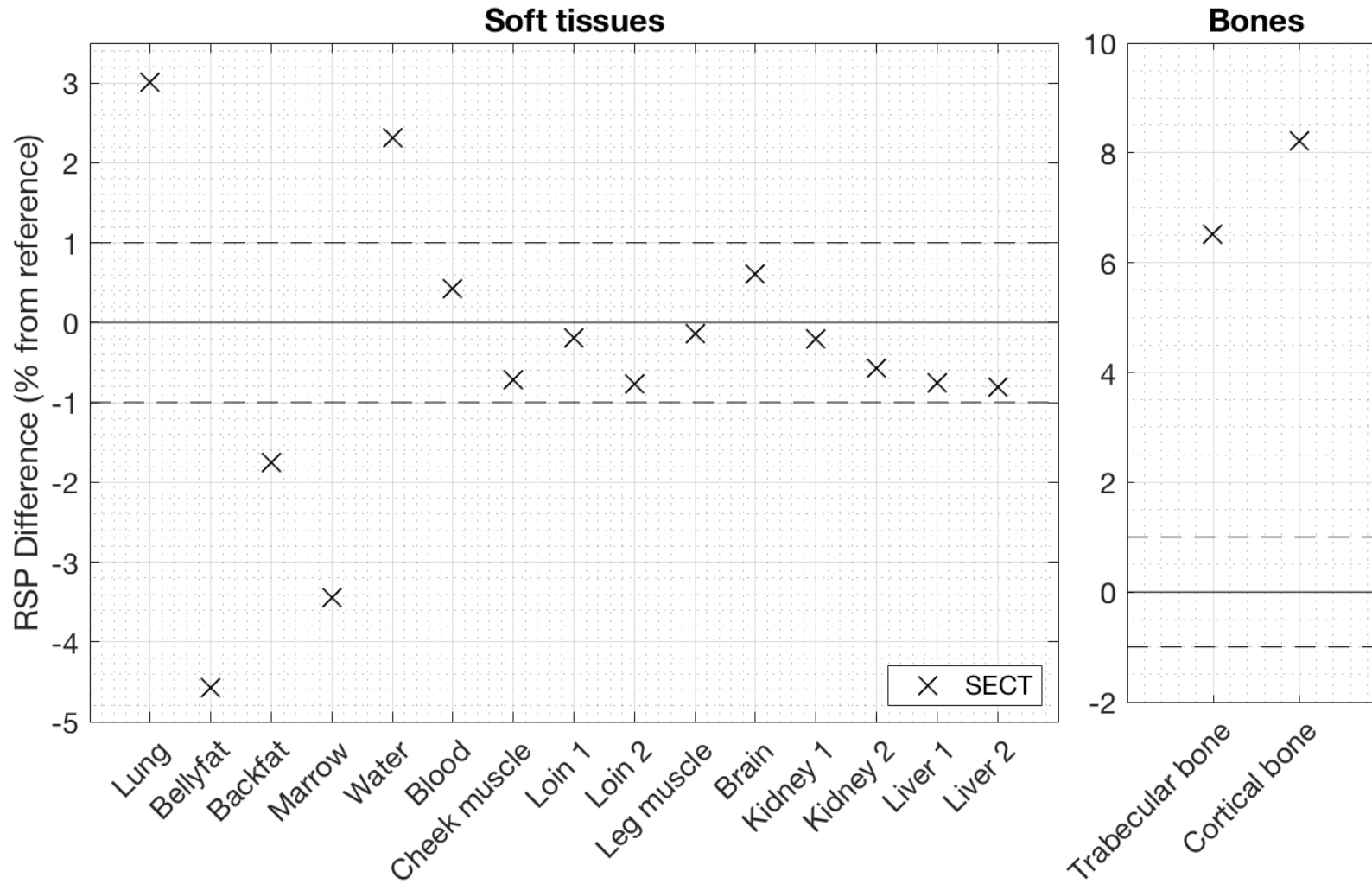
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- Calibration phantom: Gammex RMI 467 electron density phantom
- Measure average RSP per sample in VOIs:  $RSP_{DECT}$

$$\Delta RSP(\%) = \frac{RSP_{DECT} - RSP_{ref}}{RSP_{ref}} * 100$$

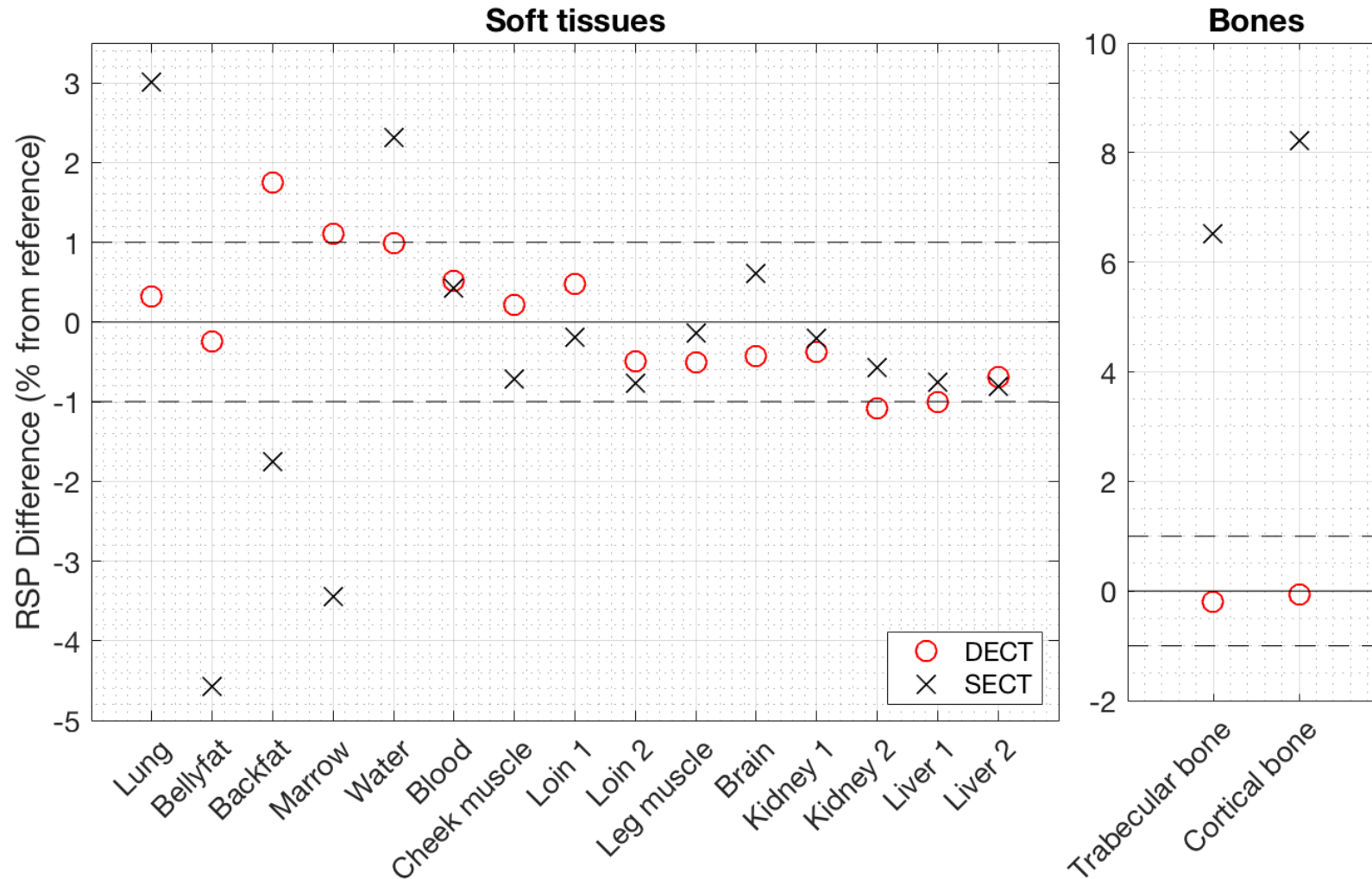


# SECT-estimated RSP errors



$$\Delta RSP(\%) = \frac{RSP_{SECT} - RSP_{ref}}{RSP_{ref}} * 100$$

# Photon CT estimated RSP errors



$$\Delta RSP(\%) = \frac{RSP_{SECT} - RSP_{ref}}{RSP_{ref}} * 100$$

$$\Delta RSP(\%) = \frac{RSP_{DECT} - RSP_{ref}}{RSP_{ref}} * 100$$

# DECT-estimated RSP errors

Tissue	RSP <sub>ref</sub>	% Difference	
		SECT	DECT
Lung	0.90 ± 0.04	3.01 ± 11.06	0.32 ± 10.87
Belly fat	1.00 ± 0.00	-4.57 ± 3.35*	-0.24 ± 4.24
Back fat	0.97 ± 0.01	-1.75 ± 1.97	1.74 ± 3.30 <sup>†</sup>
Marrow	0.93 ± 0.02	-3.44 ± 5.78	1.11 ± 7.01
Water	1.00 ± 0.00	2.32 ± 0.31	0.99 ± 2.31
Blood	1.05 ± 0.00	0.43 ± 0.29	0.51 ± 1.94
Cheek muscle	1.05 ± 0.01	-0.72 ± 1.49	0.22 ± 2.96
Loin 1	1.06 ± 0.00	-0.20 ± 2.18	0.47 ± 3.57
Loin 2	1.06 ± 0.00	-0.77 ± 1.51	-0.49 ± 2.78
Leg muscle	1.05 ± 0.01	-0.14 ± 1.68	-0.50 ± 2.78
Brain	1.04 ± 0.00	0.61 ± 0.83	-0.43 ± 2.13
Kidney 1	1.05 ± 0.00	-0.20 ± 3.25	-0.37 ± 3.82
Kidney 2	1.04 ± 0.01	-0.58 ± 4.90	-1.09 ± 6.27 *
Liver 1	1.06 ± 0.00	-0.75 ± 4.61	-1.01 ± 5.23
Liver 2	1.06 ± 0.00	-0.80 ± 6.31	-0.69 ± 6.90
Trabecular bone	1.19 ± 0.06	6.52 ± 6.86	-0.19 ± 6.61
Cortical bone	1.78 ± 0.03	8.22 ± 5.06 <sup>†</sup>	-0.07 ± 4.87
Mean error	–	<b>0.42 ± 4.55</b>	<b>-0.02 ± 5.11</b>
Mean absolute error	–	<b>2.06</b>	<b>0.61</b>
Root mean square error	–	<b>3.10</b>	<b>0.75</b>

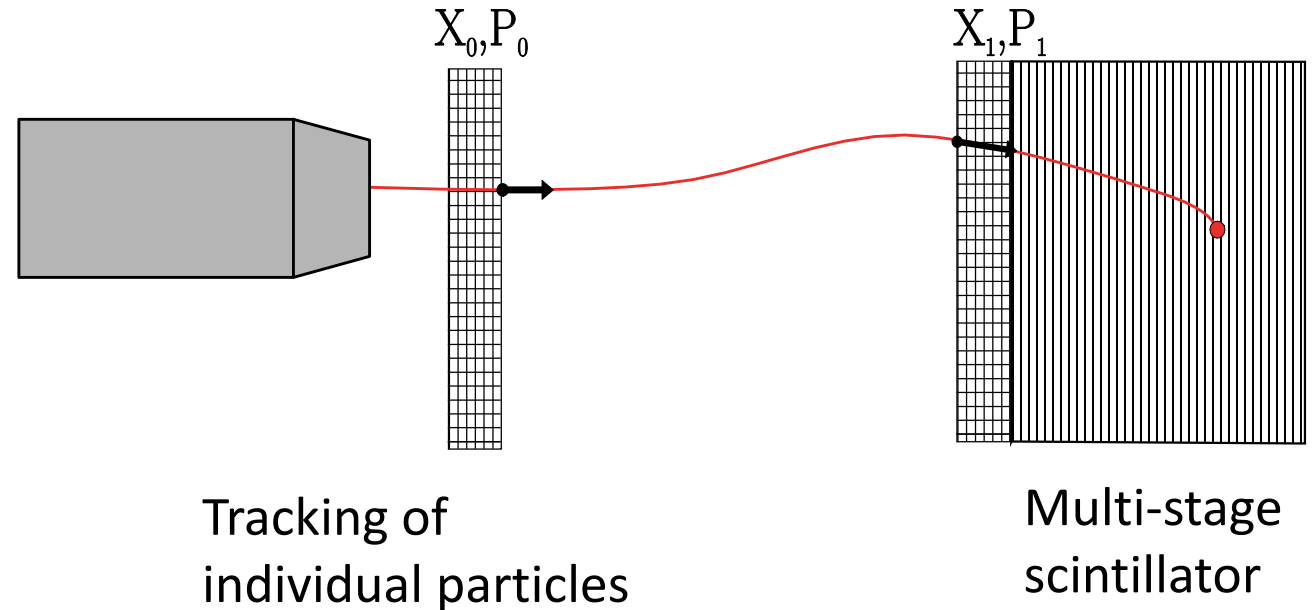
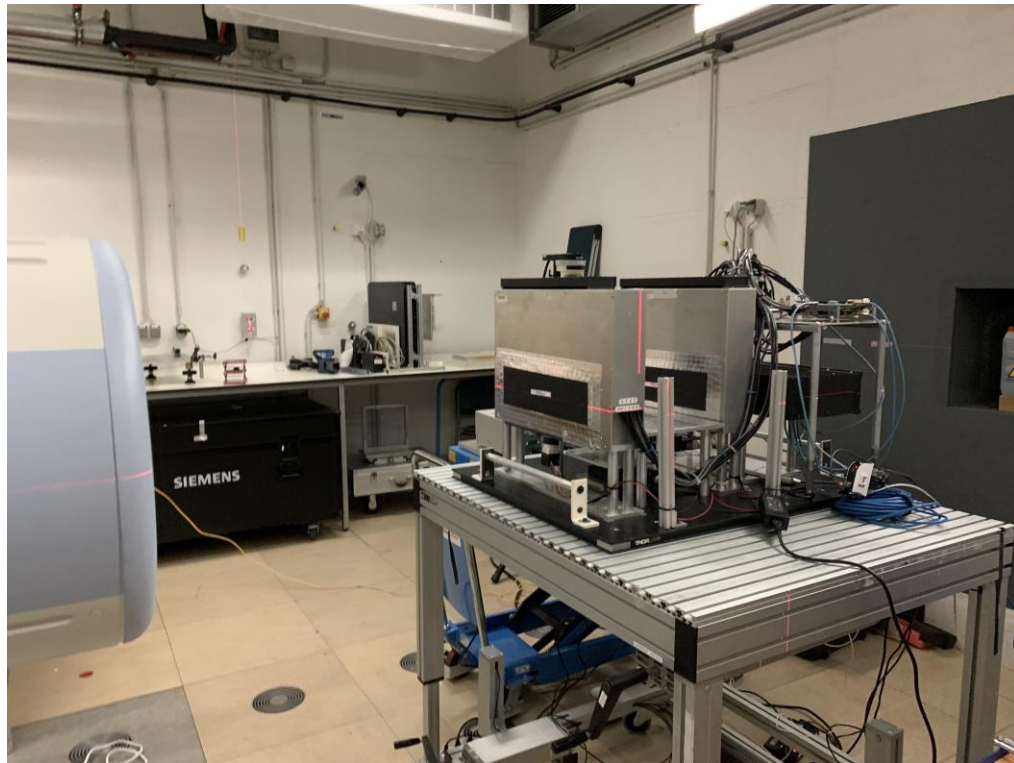
$$\Delta\text{RSP}(\%) = \frac{\text{RSP}_{\text{DECT}} - \text{RSP}_{\text{ref}}}{\text{RSP}_{\text{ref}}} * 100$$



# Which imaging modality provides the highest RSP accuracy?

- 1) Design a **tissue phantom**, collect and prepare tissue samples.
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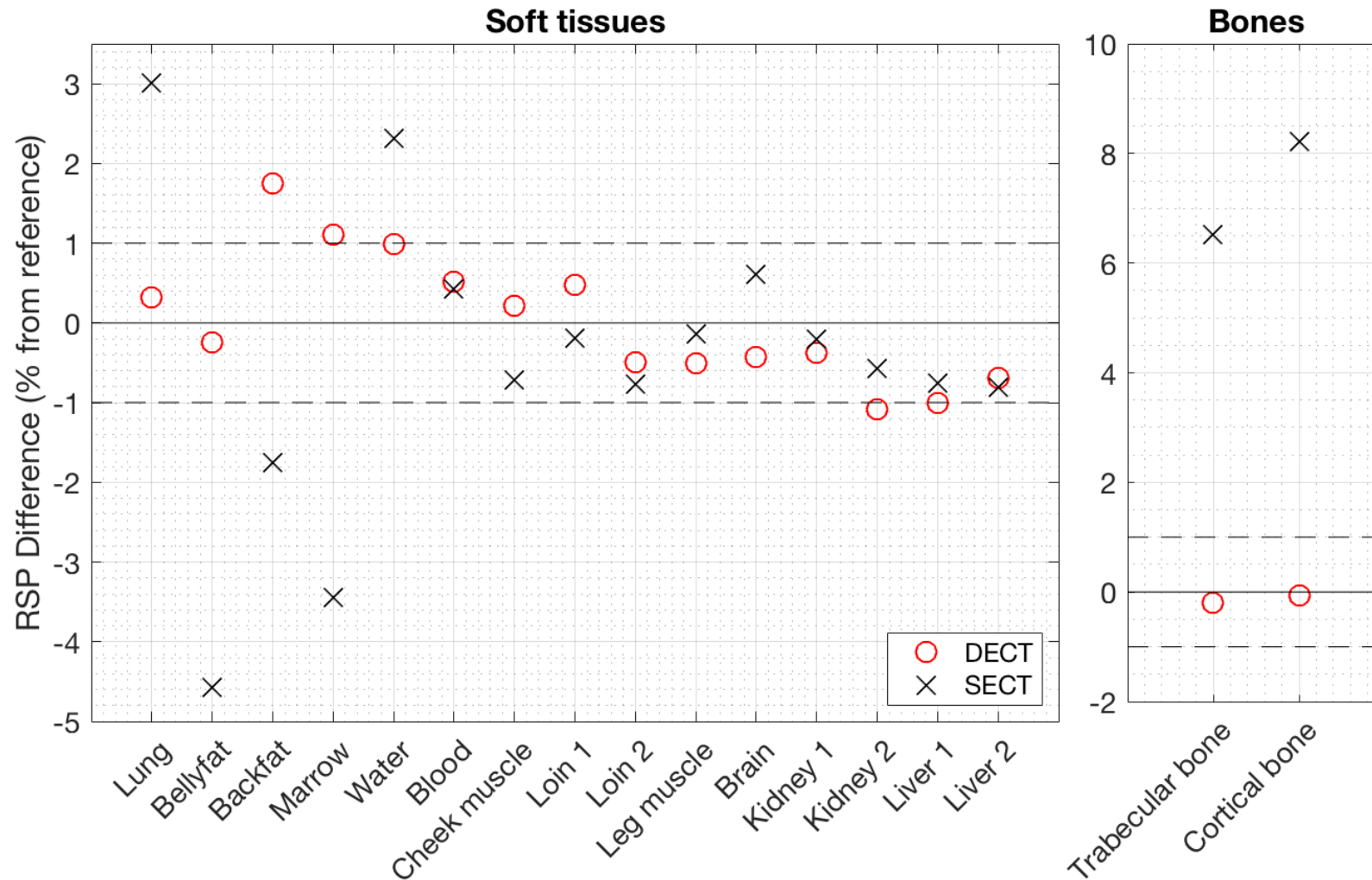
# Proton and helium CT collection



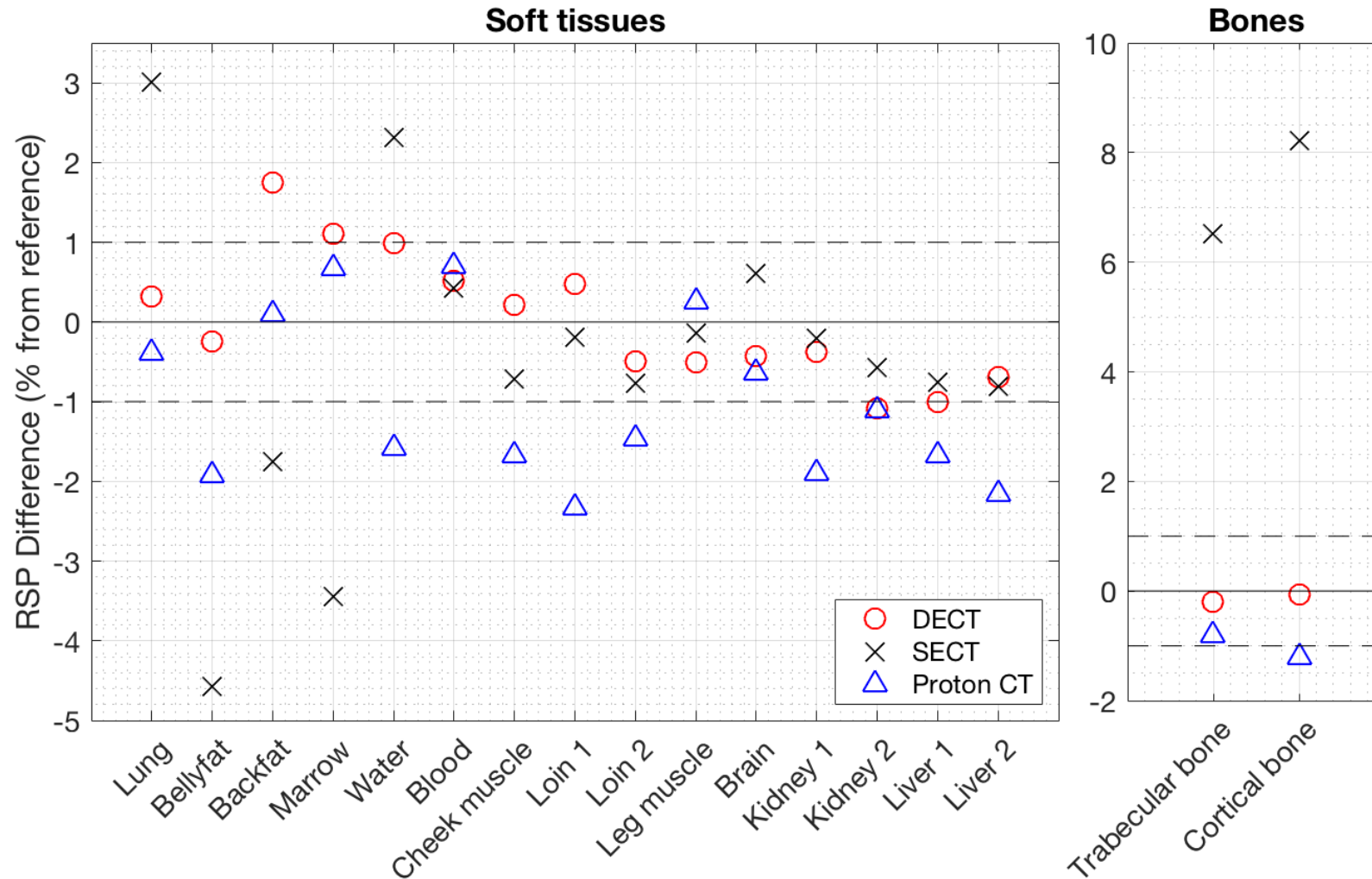
- Particle CT directly samples energy loss
- Image blur due to non-linear path of protons
- Helium ions are investigated

$$\Delta\text{RSP}(\%) = \frac{\text{RSP}_{\text{pCT}} - \text{RSP}_{\text{ref}}}{\text{RSP}_{\text{ref}}} * 100$$

# Photon CT estimated RSP errors



# Photon CT vs. Proton CT

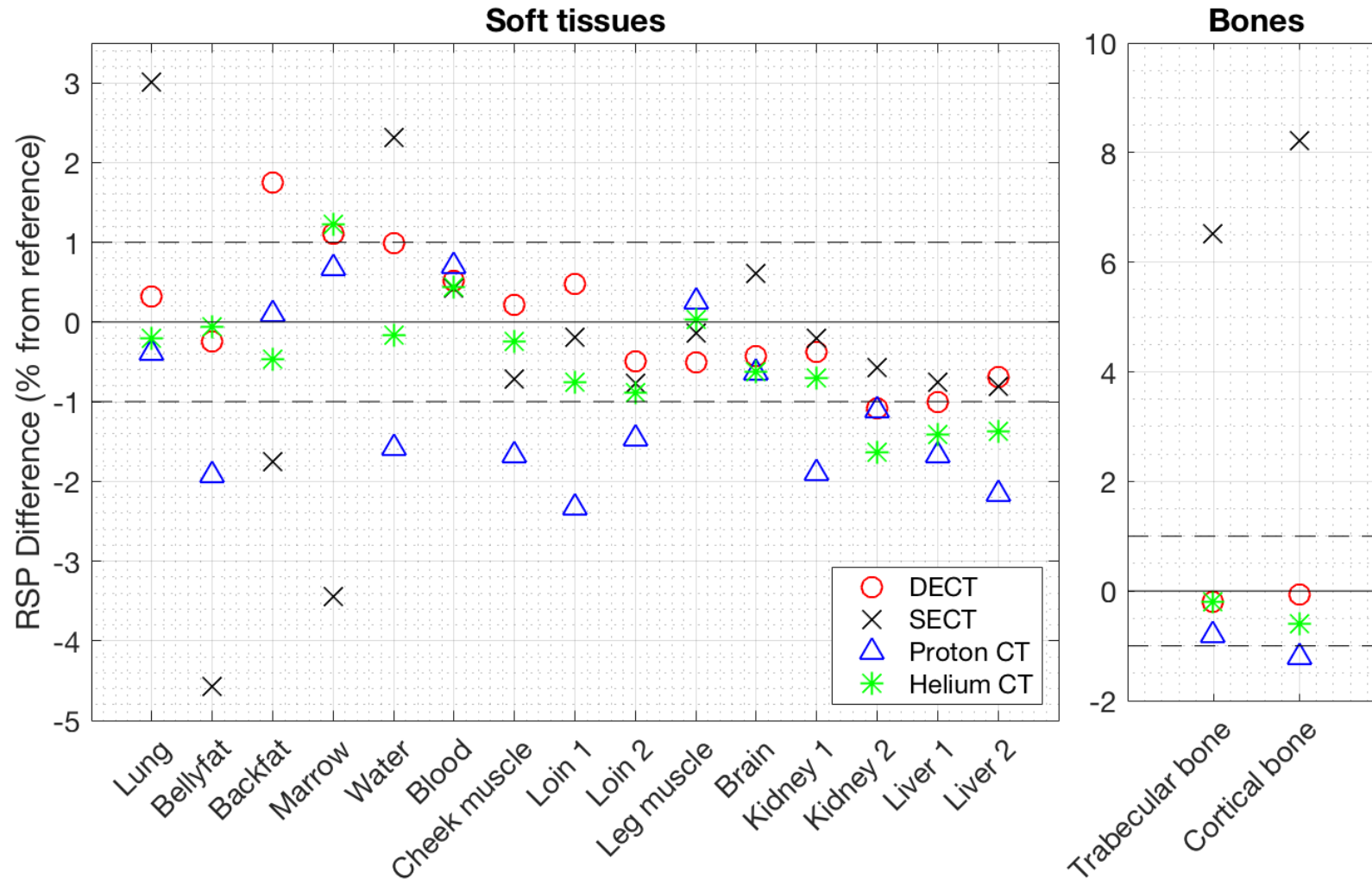




# Proton CT estimated RSP errors

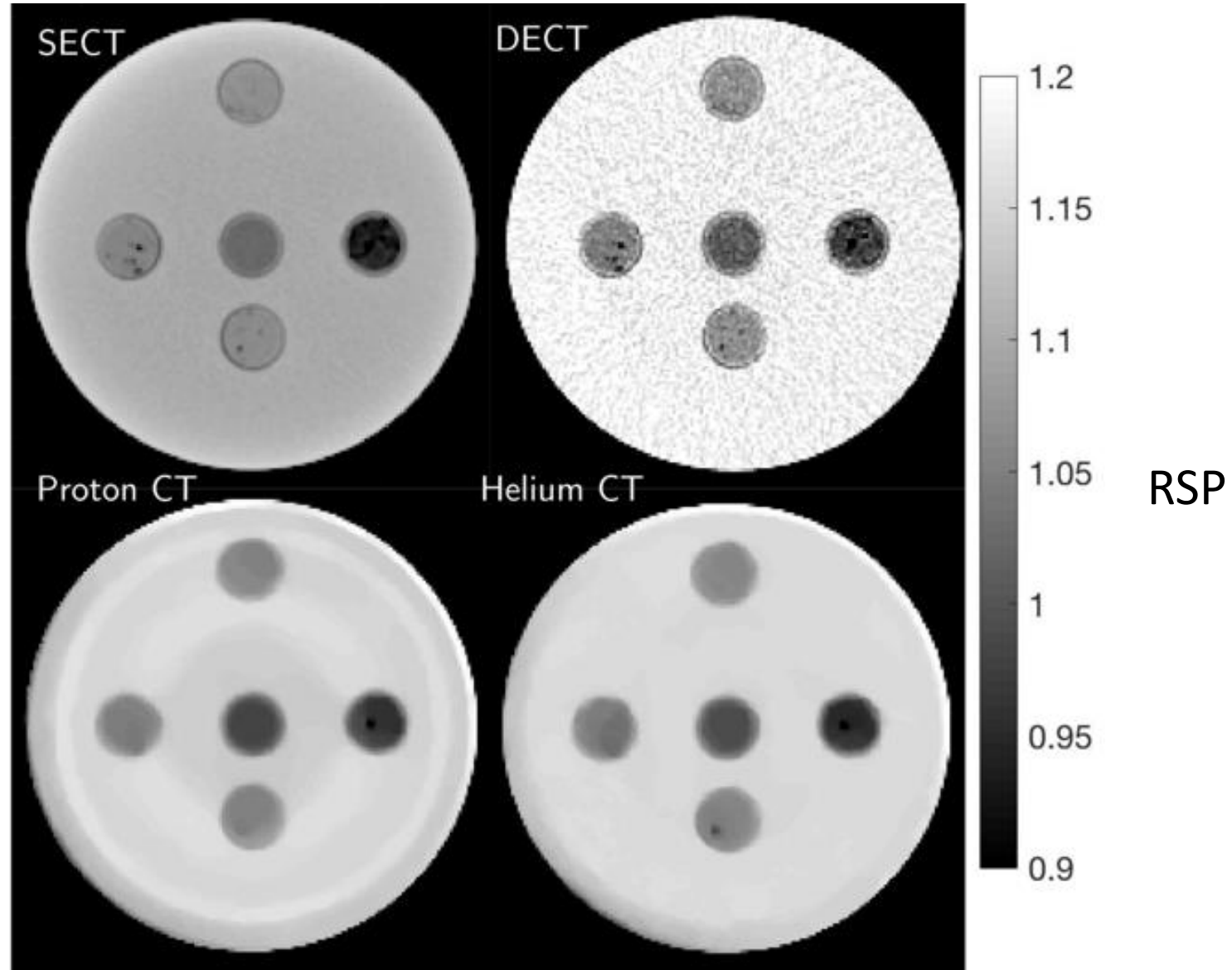
Tissue	RSP <sub>ref</sub>	% Difference to reference		
		SECT	DECT	proton CT
Lung	0.90 ± 0.04	3.01 ± 11.06	0.32 ± 10.87	-0.39 ± 6.89
Belly fat	1.00 ± 0.00	-4.57 ± 3.35*	-0.24 ± 4.24	-1.92 ± 0.83
Back fat	0.97 ± 0.01	-1.75 ± 1.97	1.74 ± 3.30 <sup>†</sup>	0.10 ± 0.95
Marrow	0.93 ± 0.02	-3.44 ± 5.78	1.11 ± 7.01	0.68 ± 4.23
Water	1.00 ± 0.00	2.32 ± 0.31	0.99 ± 2.31	-1.59 ± 0.21
Blood	1.05 ± 0.00	0.43 ± 0.29	0.51 ± 1.94	0.70 ± 0.29 <sup>†</sup>
Cheek muscle	1.05 ± 0.01	-0.72 ± 1.49	0.22 ± 2.96	-1.68 ± 1.12
Loin 1	1.06 ± 0.00	-0.20 ± 2.18	0.47 ± 3.57	-2.33 ± 1.01*
Loin 2	1.06 ± 0.00	-0.77 ± 1.51	-0.49 ± 2.78	-1.47 ± 0.72
Leg muscle	1.05 ± 0.01	-0.14 ± 1.68	-0.50 ± 2.78	0.26 ± 0.95
Brain	1.04 ± 0.00	0.61 ± 0.83	-0.43 ± 2.13	-0.64 ± 0.31
Kidney 1	1.05 ± 0.00	-0.20 ± 3.25	-0.37 ± 3.82	-1.90 ± 0.85
Kidney 2	1.04 ± 0.01	-0.58 ± 4.90	-1.09 ± 6.27 *	-1.12 ± 2.12
Liver 1	1.06 ± 0.00	-0.75 ± 4.61	-1.01 ± 5.23	-1.68 ± 1.50
Liver 2	1.06 ± 0.00	-0.80 ± 6.31	-0.69 ± 6.90	-2.16 ± 5.32
Trabecular bone	1.19 ± 0.06	6.52 ± 6.86	-0.19 ± 6.61	-0.82 ± 5.83
Cortical bone	1.78 ± 0.03	8.22 ± 5.06 <sup>†</sup>	-0.07 ± 4.87	-1.22 ± 1.58
Mean error	–	<b>0.42 ± 4.55</b>	<b>-0.02 ± 5.11</b>	<b>-1.01 ± 2.90</b>
Mean absolute error	–	<b>2.06</b>	<b>0.61</b>	<b>1.21</b>
Root mean square error	–	<b>3.10</b>	<b>0.75</b>	<b>1.39</b>

# Photon vs. Particle CT



# Particle CT

Tissue	RSP <sub>ref</sub>	% Difference to reference			
		SECT	DECT	proton CT	Helium CT
Lung	0.90 ± 0.04	3.01 ± 11.06	0.32 ± 10.87	-0.39 ± 6.89	-0.21 ± 7.13
Belly fat	1.00 ± 0.00	-4.57 ± 3.35*	-0.24 ± 4.24	-1.92 ± 0.83	-0.07 ± 1.32
Back fat	0.97 ± 0.01	-1.75 ± 1.97	1.74 ± 3.30 <sup>†</sup>	0.10 ± 0.95	-0.47 ± 0.98
Marrow	0.93 ± 0.02	-3.44 ± 5.78	1.11 ± 7.01	0.68 ± 4.23	1.23 ± 3.74 <sup>†</sup>
Water	1.00 ± 0.00	2.32 ± 0.31	0.99 ± 2.31	-1.59 ± 0.21	-0.16 ± 0.25
Blood	1.05 ± 0.00	0.43 ± 0.29	0.51 ± 1.94	0.70 ± 0.29 <sup>†</sup>	0.43 ± 0.18
Cheek muscle	1.05 ± 0.01	-0.72 ± 1.49	0.22 ± 2.96	-1.68 ± 1.12	-0.24 ± 1.17
Loin 1	1.06 ± 0.00	-0.20 ± 2.18	0.47 ± 3.57	-2.33 ± 1.01*	-0.75 ± 1.38
Loin 2	1.06 ± 0.00	-0.77 ± 1.51	-0.49 ± 2.78	-1.47 ± 0.72	-0.89 ± 0.98
Leg muscle	1.05 ± 0.01	-0.14 ± 1.68	-0.50 ± 2.78	0.26 ± 0.95	0.03 ± 1.11
Brain	1.04 ± 0.00	0.61 ± 0.83	-0.43 ± 2.13	-0.64 ± 0.31	-0.63 ± 0.48
Kidney 1	1.05 ± 0.00	-0.20 ± 3.25	-0.37 ± 3.82	-1.90 ± 0.85	-0.70 ± 1.07
Kidney 2	1.04 ± 0.01	-0.58 ± 4.90	-1.09 ± 6.27 *	-1.12 ± 2.12	-1.64 ± 3.34*
Liver 1	1.06 ± 0.00	-0.75 ± 4.61	-1.01 ± 5.23	-1.68 ± 1.50	-1.41 ± 1.40
Liver 2	1.06 ± 0.00	-0.80 ± 6.31	-0.69 ± 6.90	-2.16 ± 5.32	-1.37 ± 5.34
Trabecular bone	1.19 ± 0.06	6.52 ± 6.86	-0.19 ± 6.61	-0.82 ± 5.83	-0.19 ± 6.22
Cortical bone	1.78 ± 0.03	8.22 ± 5.06 <sup>†</sup>	-0.07 ± 4.87	-1.22 ± 1.58	-0.60 ± 2.02
Mean error	–	<b>0.42 ± 4.55</b>	<b>-0.02 ± 5.11</b>	<b>-1.01 ± 2.90</b>	<b>-0.50 ± 3.06</b>
Mean absolute error	–	<b>2.06</b>	<b>0.61</b>	<b>1.21</b>	<b>0.65</b>
Root mean square error	–	<b>3.10</b>	<b>0.75</b>	<b>1.39</b>	<b>0.81</b>



# Conclusion

- We present a first comparison of photon and particle CT for RSP estimation based on fresh tissues
- SECT is highly biased in low- and high density tissues
- DECT offers high accuracy
- Proton CT is currently limited by ring artefacts
- Helium CT provides good RSP accuracy and low noise
- Mind the maturity of systems!

# Acknowledgements

Sample preparation

Photon CT

Beam time



Particle CT equipment



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Data processing algorithms

