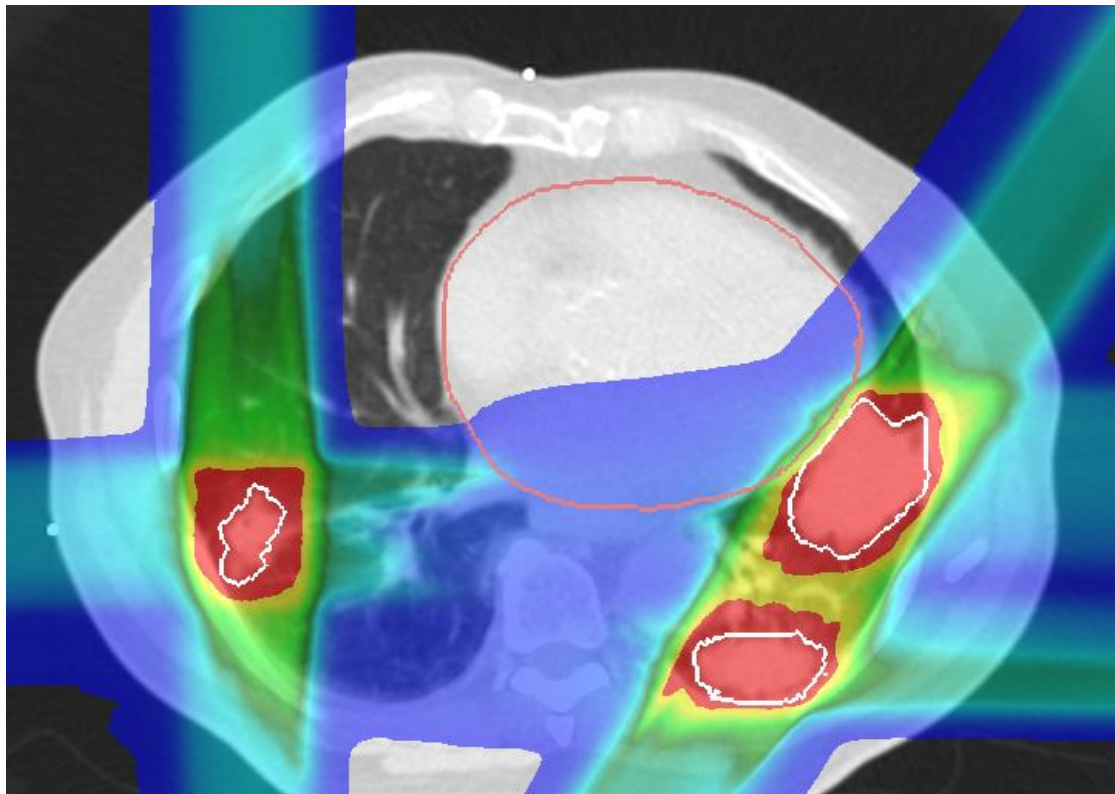


In Silico Comparison of Photons versus Carbon Ions in Single Fraction Therapy of Lung Cancer



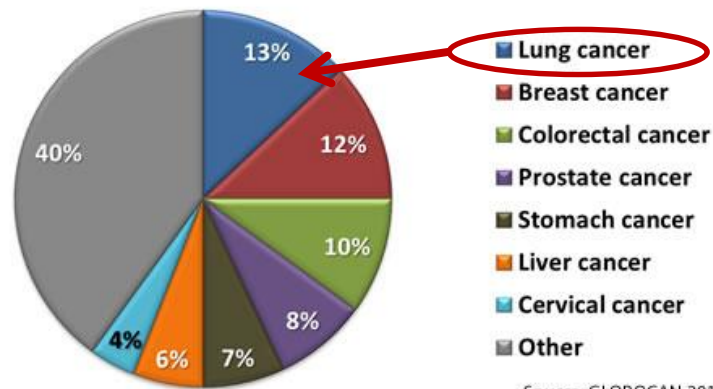
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DARMSTADT

Kristjan Anderle



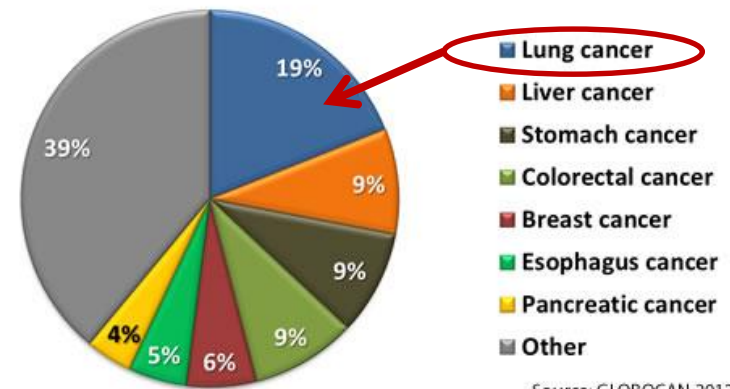
Introduction

Most Common Cancers Worldwide in 2012



Source: GLOBOCAN 2012

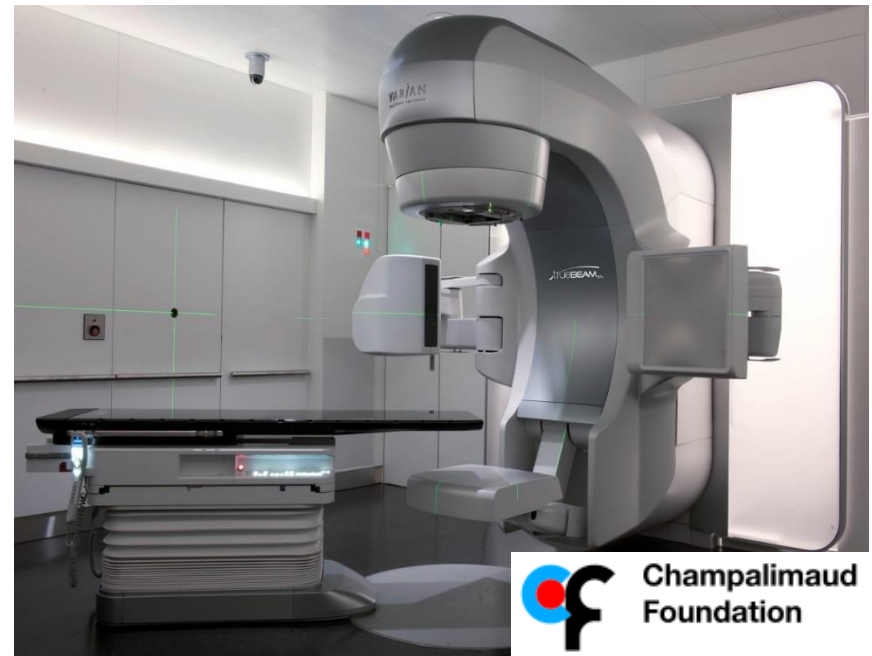
Most Common Causes of Cancer Death Worldwide in 2012



Source: GLOBOCAN 2012

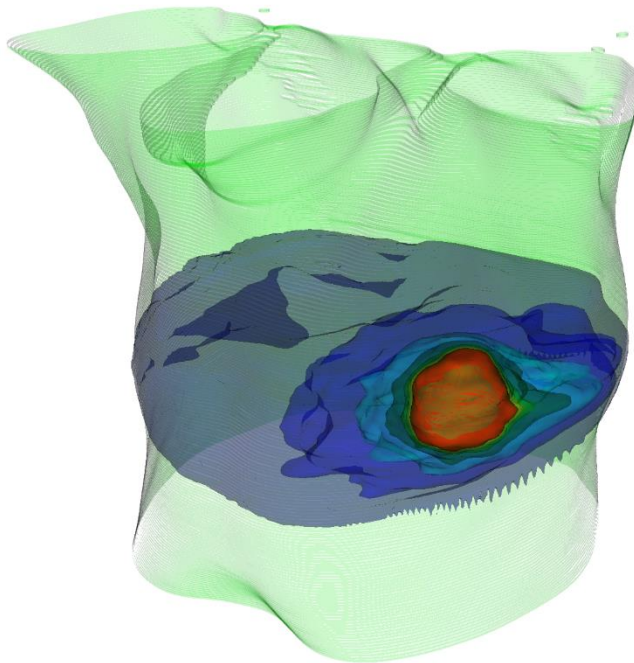
Stereotactic Body Radiation Treatment (SBRT)

- High precision treatment with photons
- Delivers high dose (up to 30 Gy) in few fractions (1-3)
- Good results for lung cancer in early stages
- Limited for:
 - Multiple tumors
 - Large tumors

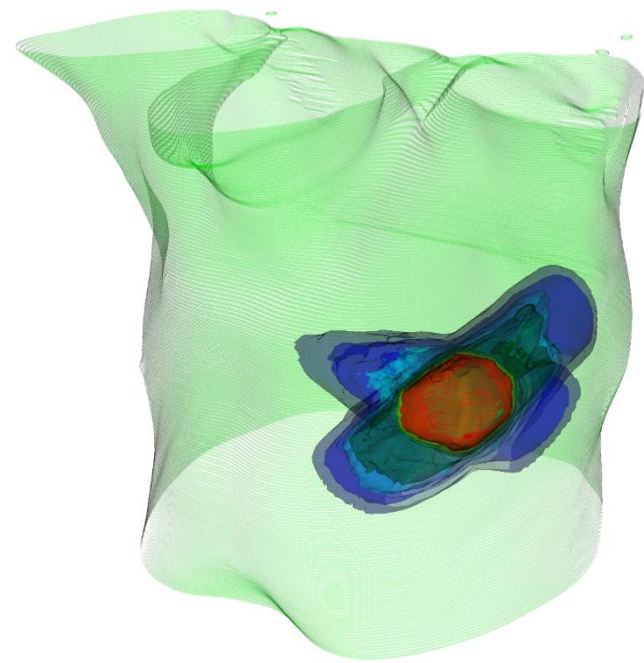


3D Treatment Plan Example

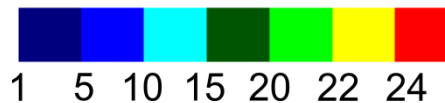
Photons



^{12}C

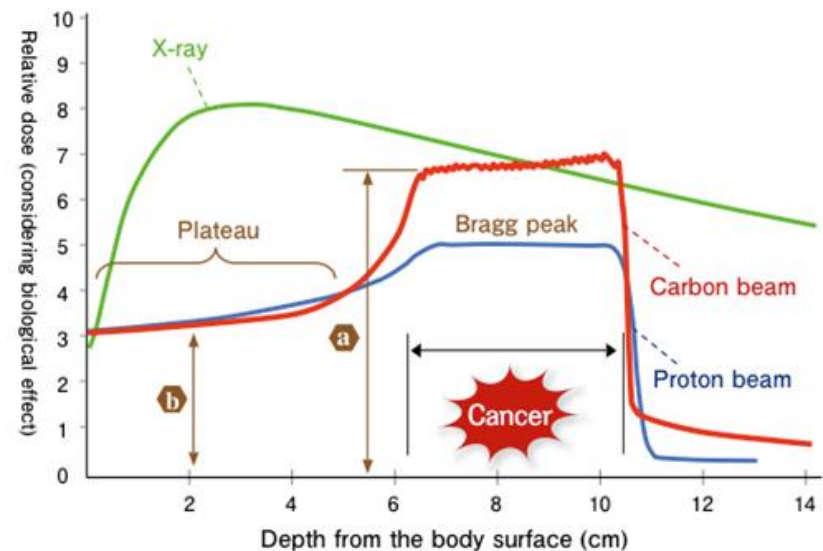


Dose (Gy)



Particle Therapy

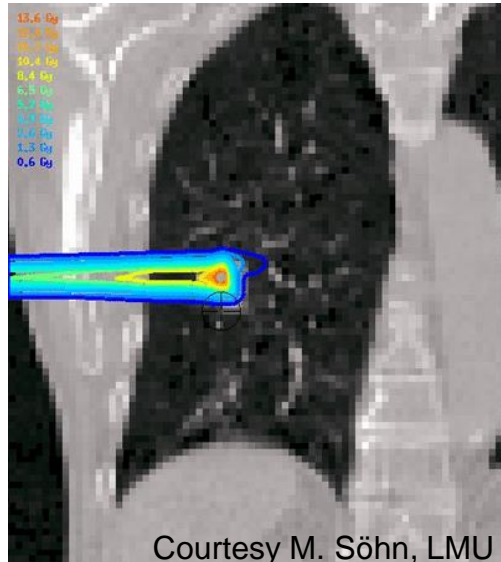
- Advantageous dose profile for radiotherapy
- Bragg peak depends on particle energy
- No or small dose tail after the Bragg peak
- Currently, mostly static targets are treated (head & neck, pelvic)



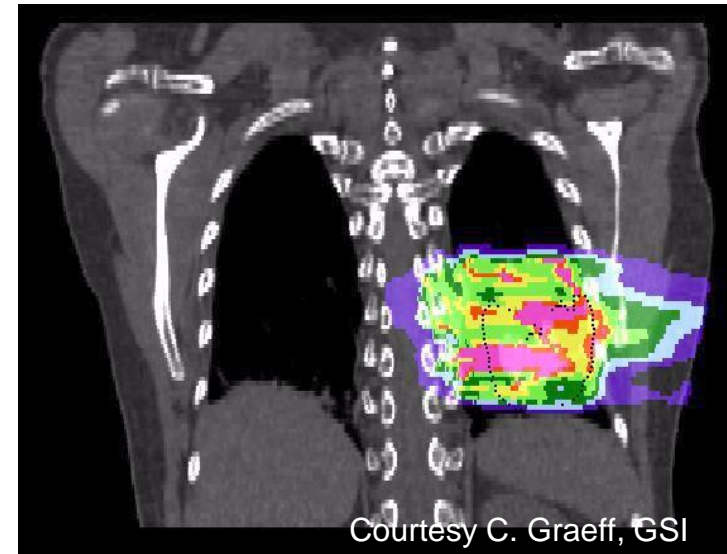
Source: NIRS, 2016

Motion Effects in Particle Therapy

Range Changes

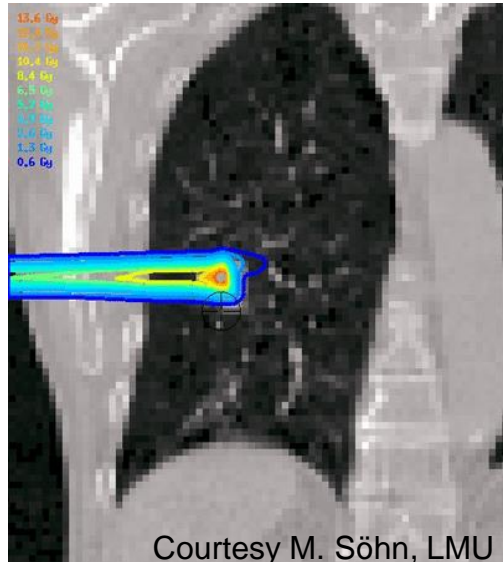


Interplay

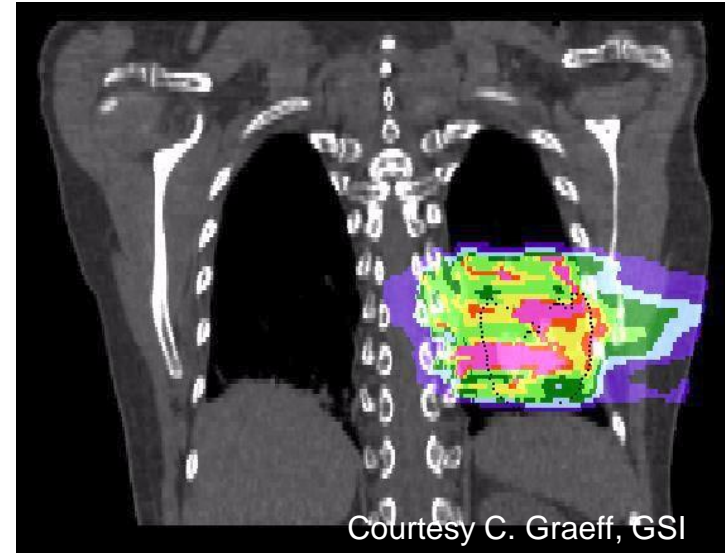


Motion Effects in Particle Therapy

Range Changes



Interplay



Compensations:

- Include range-changes in target definition (range ITV)
- Optimization in all motion states (4D-optimization)
- Rescanning: instead of delivering the whole dose at once, it is delivered in N steps

In Silico Study

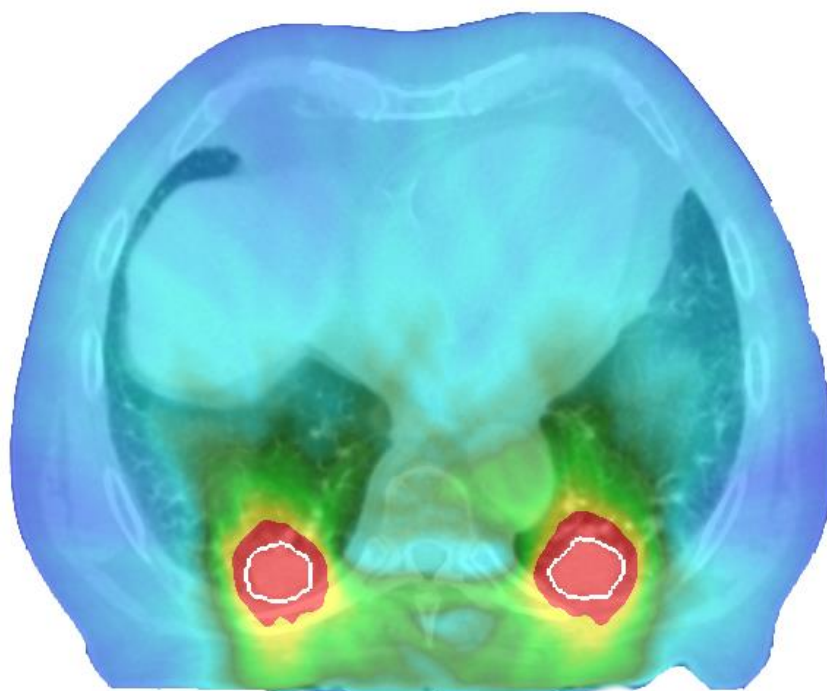
- Comparison of lung cancer treatment simulations between **SBRT** and active scanning carbon-ions (**PT**)
- 23 lung cancer patients were actually treated with SBRT
- Comparison:
 - Dose to the tumor
 - Dose to critical organs
 - Dose escalation



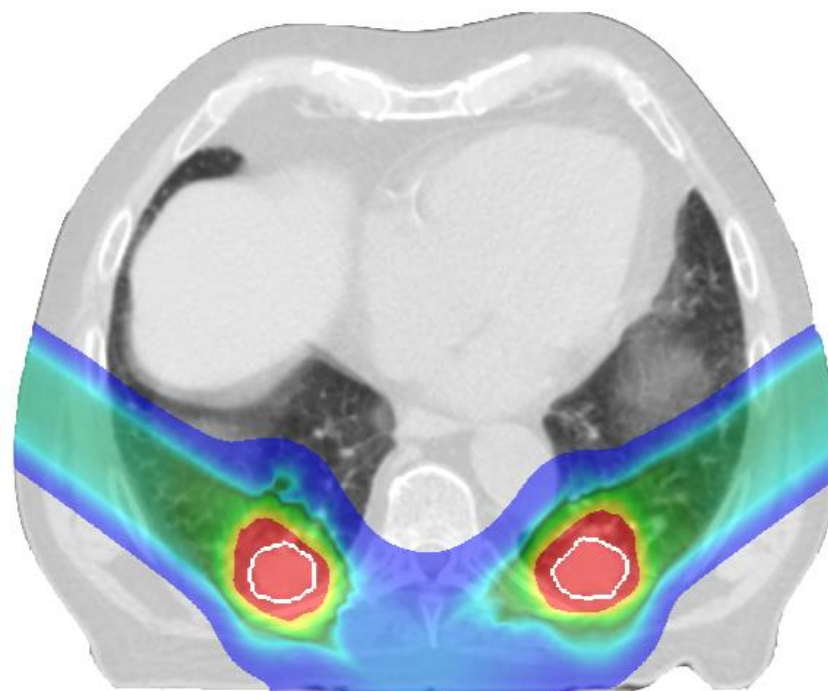
Treatment planning – Cohort I

- 19 patients, 26 tumors
- Planning objectives:
 - Single fraction of 24 Gy
 - Tumor $D_{99\%} > 100\%$
 - Critical organs limits
- Single field uniform optimization with range ITV
- 10 dose calculations per patient:
 - Without motion, static (end-inhale and end-exhale)
 - With motion, without compensation, 4D-interplay (4x)
 - With motion, with compensation, 4D-rescanning (4x)

Treatment Plan Examples



SBRT

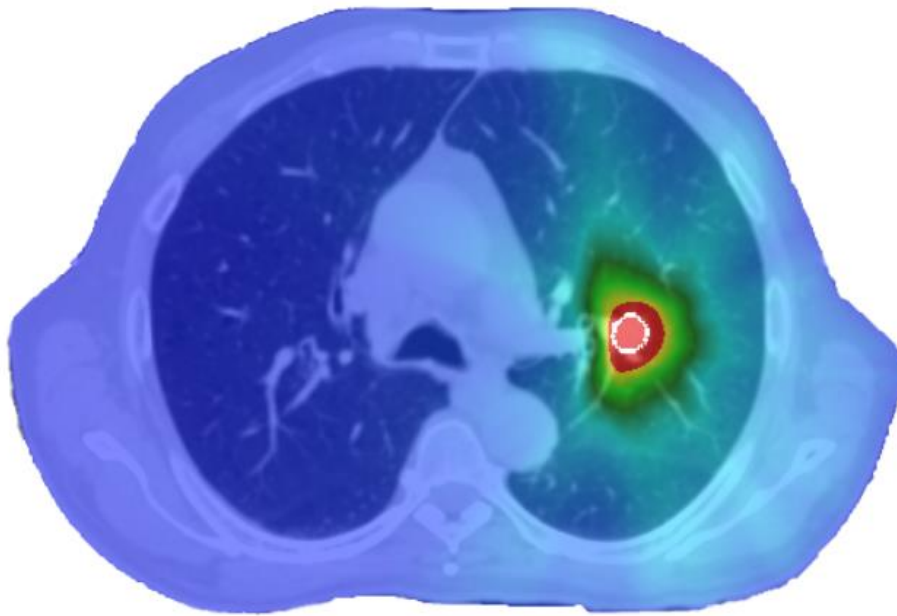


PT

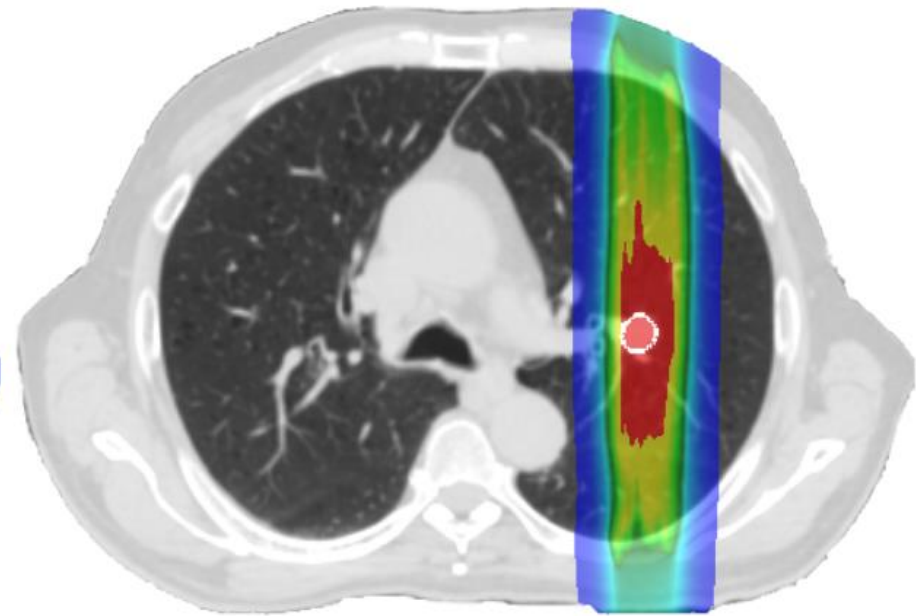
Dose (%)



Treatment Plan Examples

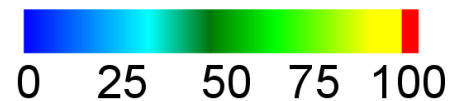


SBRT



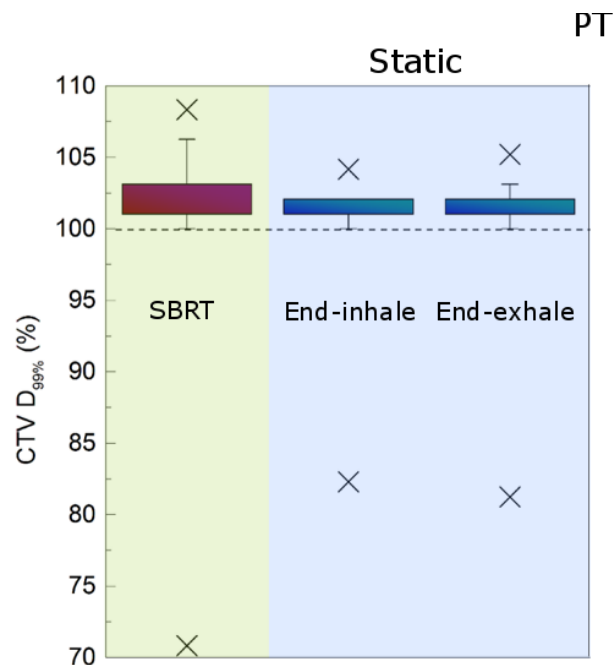
PT

Dose (%)



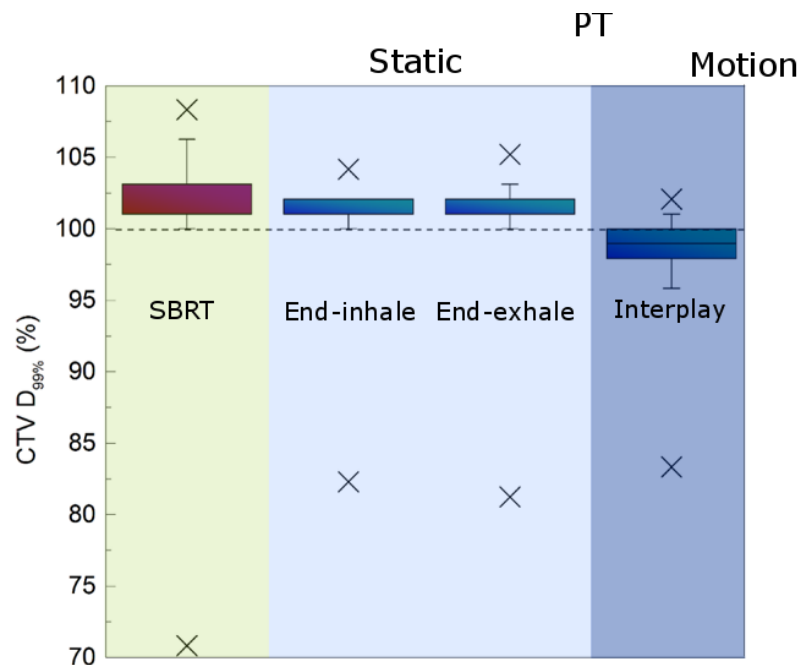
Results – Dose to Tumors

- No difference in tumor dose between SBRT and static PT (3D)



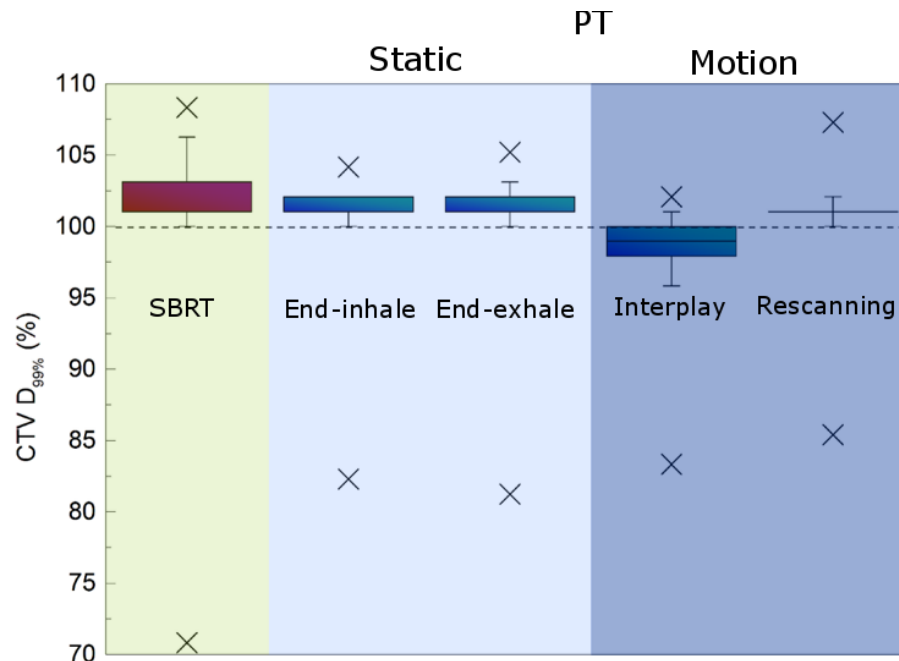
Results – Dose to Tumors

- No difference in tumor dose between SBRT and static PT (3D)
- Without motion compensation the tumor dose is too low (4D interplay)

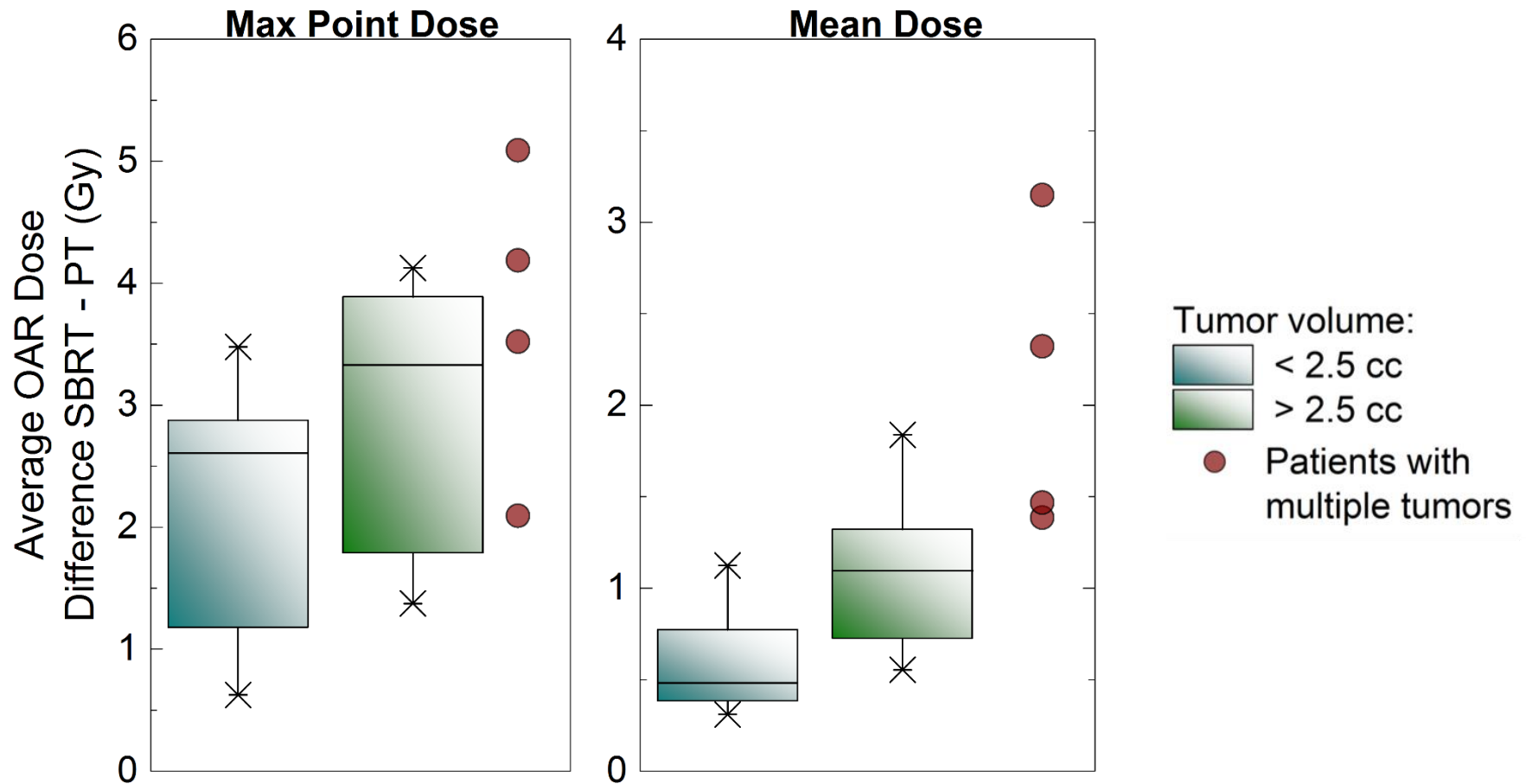


Results – Dose to Tumors

- No difference in tumor dose between SBRT and static PT (3D)
- Without motion compensation the tumor dose is too low (4D interplay)
- Rescanning mitigates tumor motion sufficiently (4D rescan)



Results - Dose to Critical Organs



Clinical Trial RTOG 0617

- A RTOG 0617 trial compared 60 Gy and 74 Gy dose to tumors
- In contrast to initial hypothesis, higher dose led to higher mortality
- The only significant difference between two groups was dose to the heart (V5)

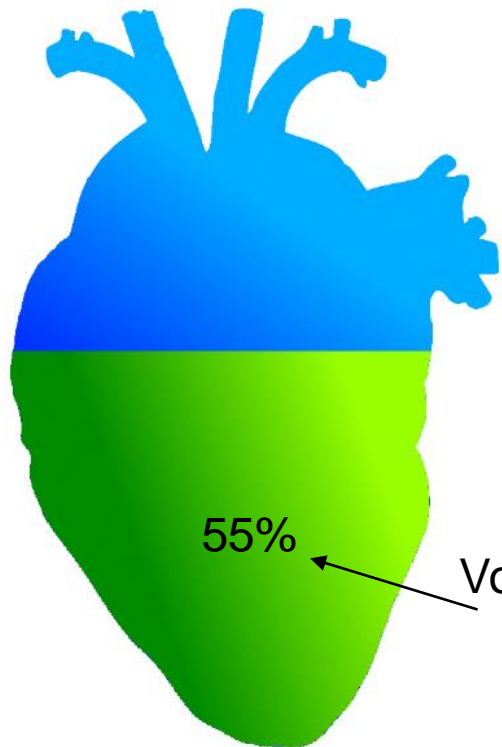
(Bradley, JD et al, Lancet Oncology, 2015)

Average Heart V5

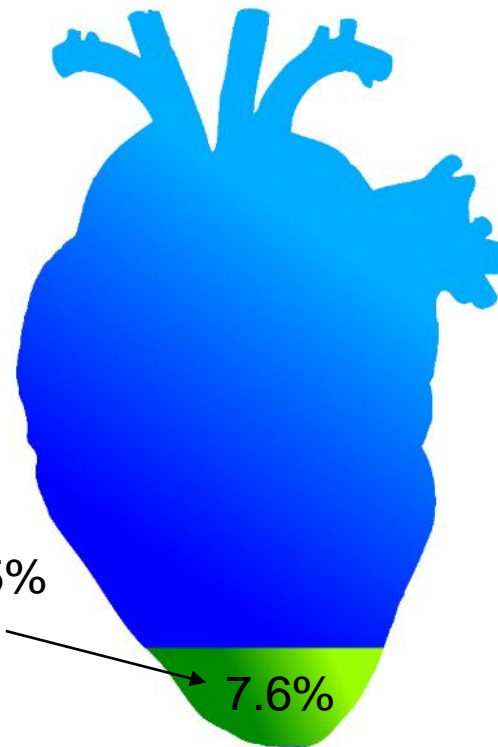


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SBRT



PT



Volume receiving 5%
dose (1.2 Gy)

Treatment planning – Cohort II

- 8 patients, 24 tumors
- Different fractionation schemes
 - 3 tumors in 2 patients received 3 x 9 Gy
 - 2 tumors received 1 x 20 Gy
 - 1 tumor received 1 x 22 Gy
 - 1 tumor received 5 x 7 Gy
 - All others 1 x 24 Gy
- Intensity modulated particle therapy optimization with rescanning
 - Range ITV
 - 4D-optimization

Treatment planning – Cohort II

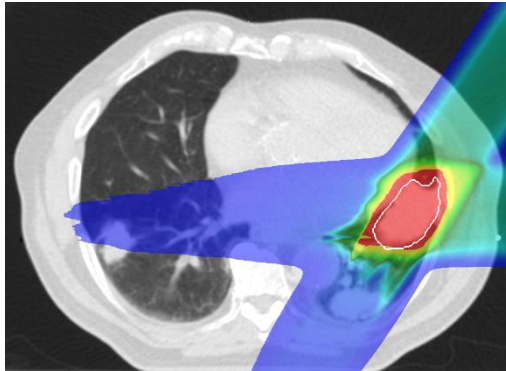
- 8 patients, 24 tumors
- Different fractionation schemes
 - 3 tumors in 2 patients received 3 x 9 Gy
 - 2 tumors received 1 x 20 Gy
 - 1 tumor received 1 x 22 Gy
 - 1 tumor received 5 x 7 Gy

} Dose escalation to 1 x 24 Gy

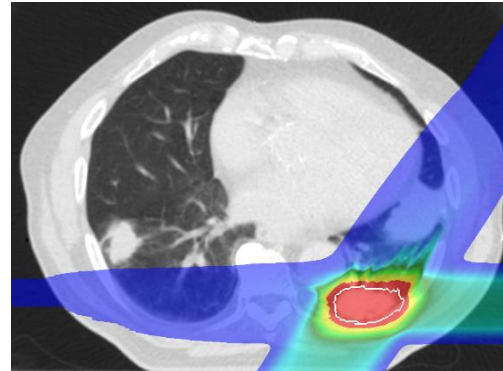
- All others 1 x 24 Gy
- Intensity modulated particle therapy optimization with rescanning
 - Range ITV
 - 4D-optimization

Intensity Modulated Particle Therapy Optimization for Multiple Targets

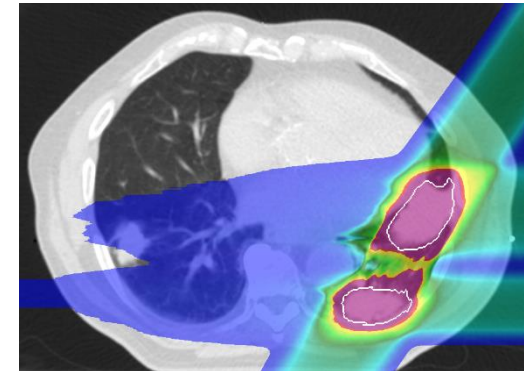
Target 1



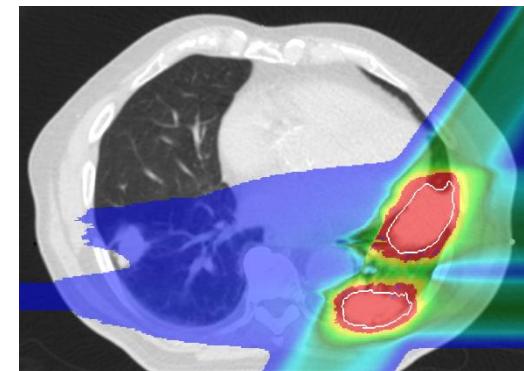
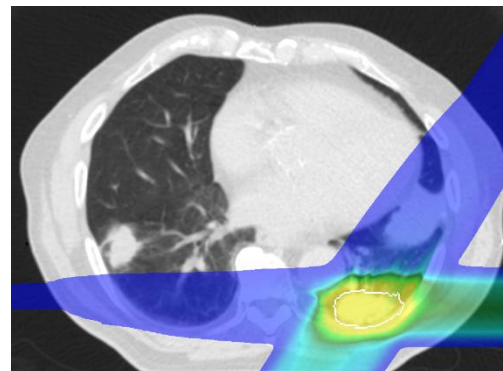
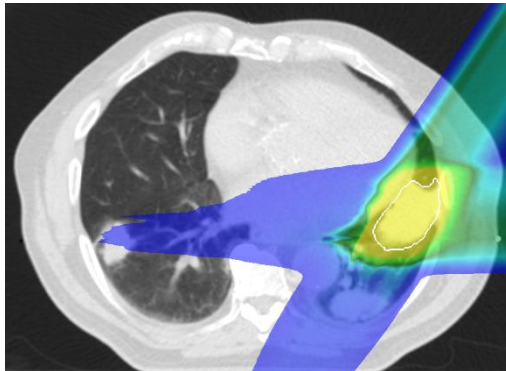
Target 2



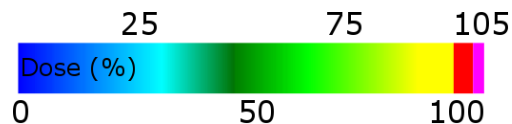
Target 1 & 2



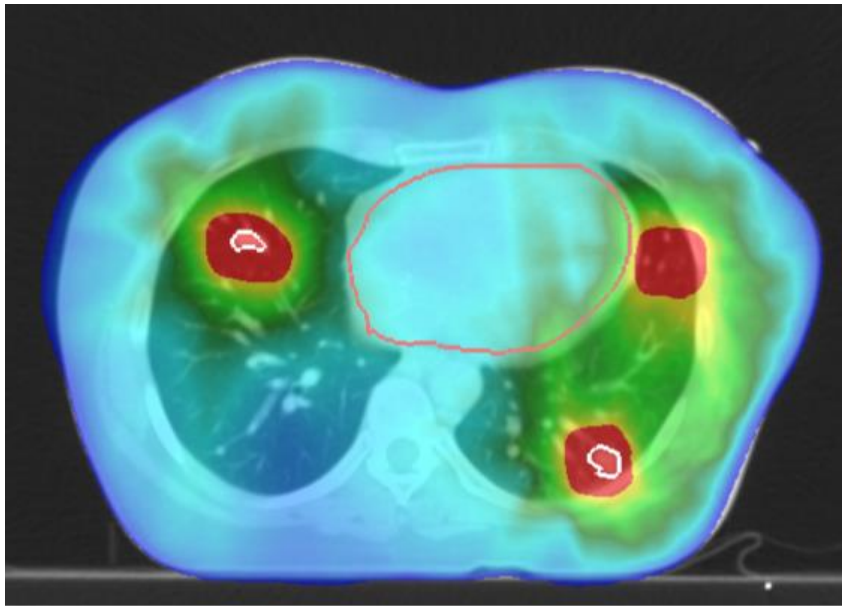
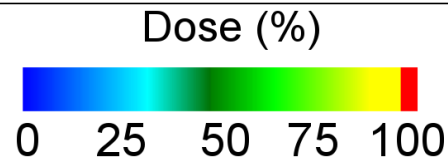
Each target
optimized
individually:



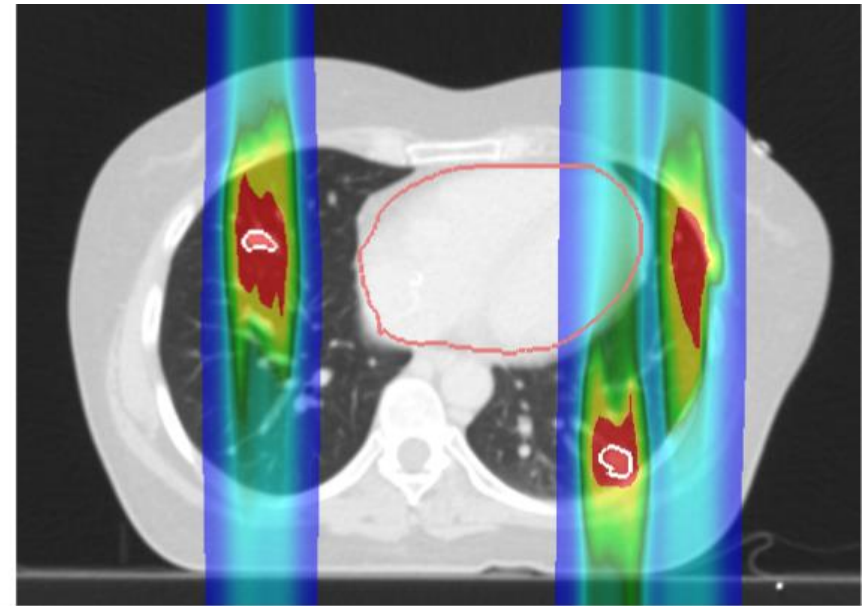
Targets
optimized
simultaneously:



Treatment Plan Examples



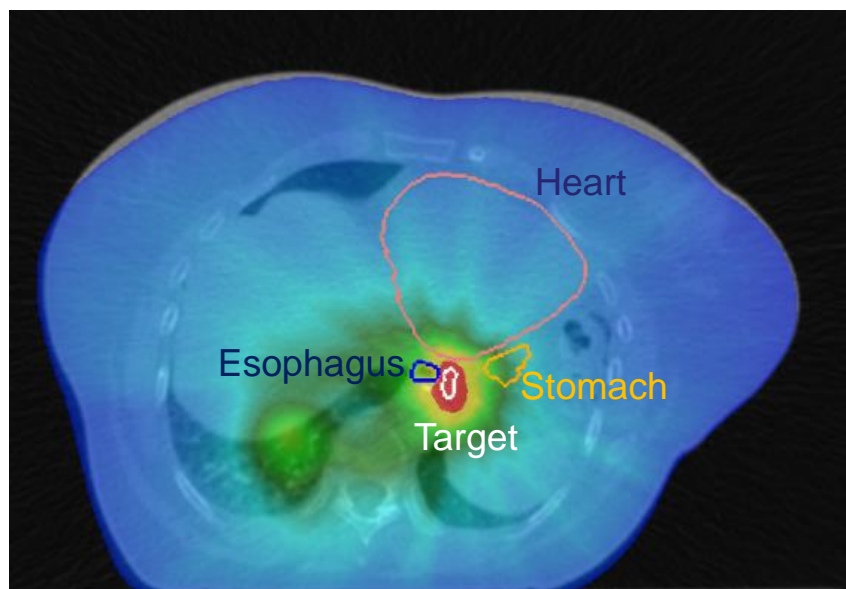
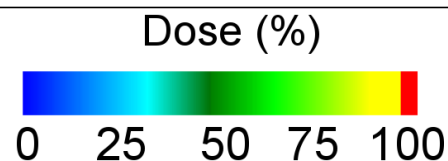
SBRT



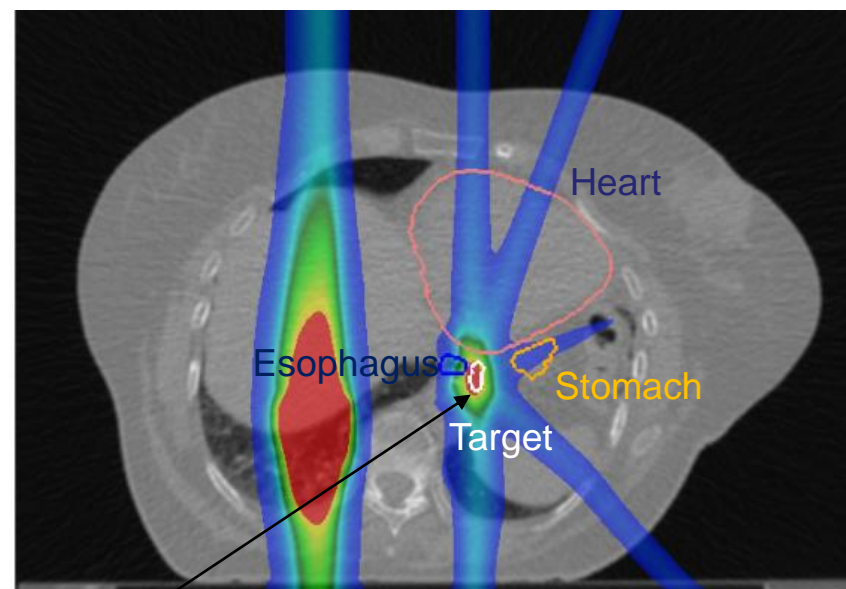
PT

All 5 targets received planned dose with SBRT and PT (CTV $D_{99\%} > 100\%$)

Treatment Plan Examples



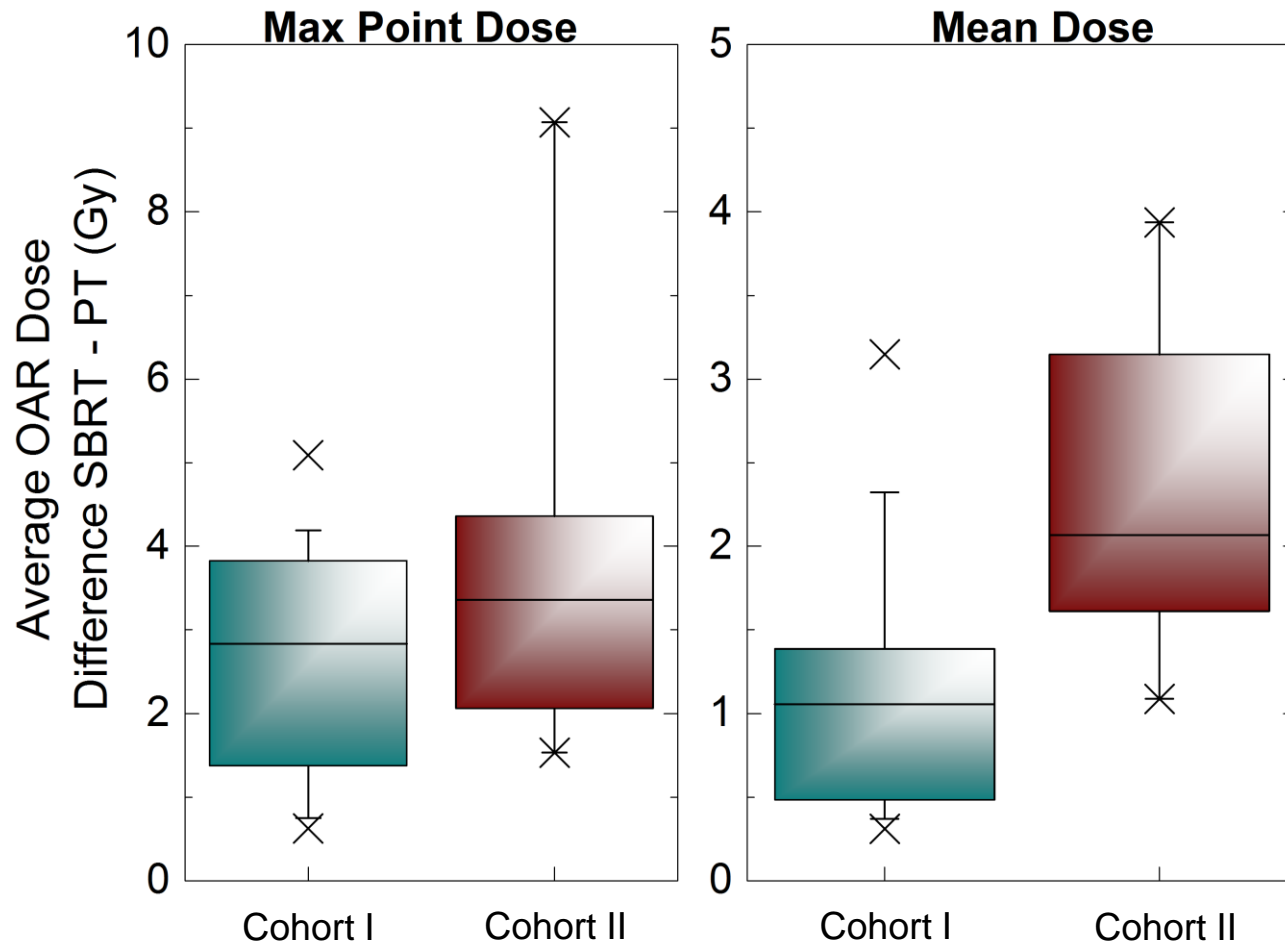
SBRT



PT

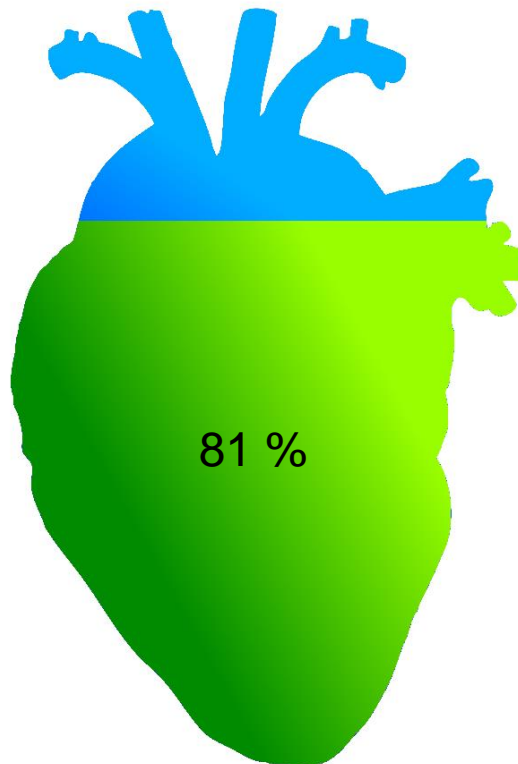
Planning dose not met with PT
(SBRT $D_{99\%} = 100\%$, PT $D_{99\%} = 75\%$)

Results – Dose to Critical Organs



Average Heart V5

SBRT

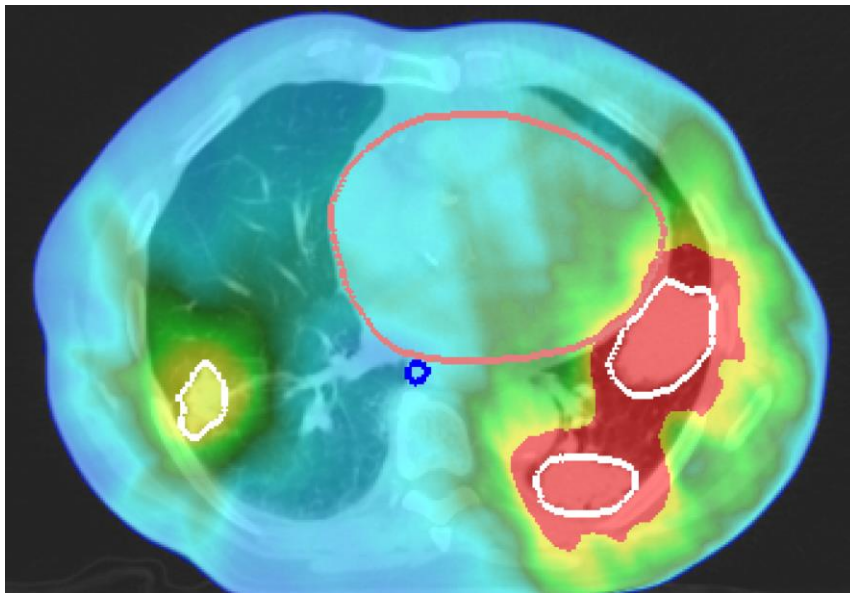


PT

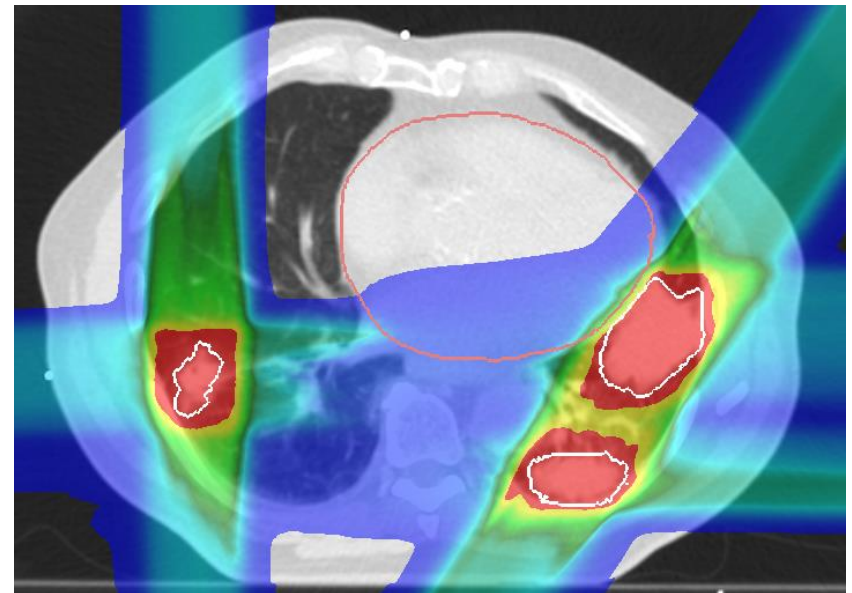


Results – Dose Escalation

Dose (%)

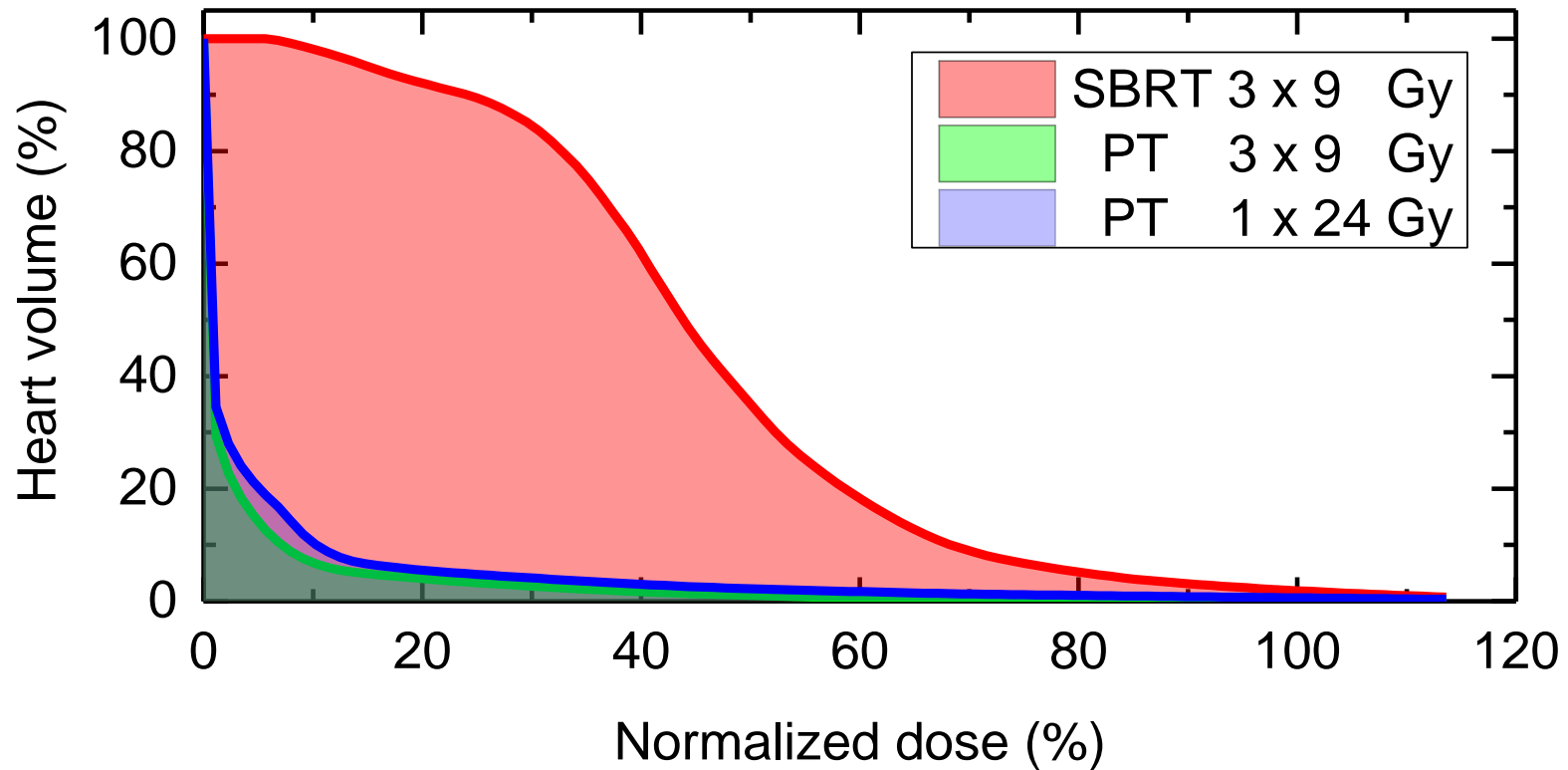


1 tumor: 1 x 22 Gy
2 tumors: 1 x 21 Gy
2 tumors: 3 x 9 Gy



5 tumors: 1 x 24 Gy

Heart Dose Volume Histogram



Conclusions

- PT delivers the same dose to tumors as SBRT, while depositing significantly less dose to normal tissue
- PT could deliver full ablative dose in single fraction, where SBRT could not
- Patient selection for PT:
 - ✓ Multiple targets
 - ✓ Large tumors
 - x Small tumors with large motions

Outlook

- First lung cancer patients treated with active scanning carbon-ions at NIRS, Chiba (Japan)
- As part of a LOEWE grant, results from this study will be used for actual lung cancer treatment at MIT – especially the multiple target case.



LOEWE

Exzellente Forschung für
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