



# Organ motion: clinical practice

Alessandro Vai

*Medical Physics department,*

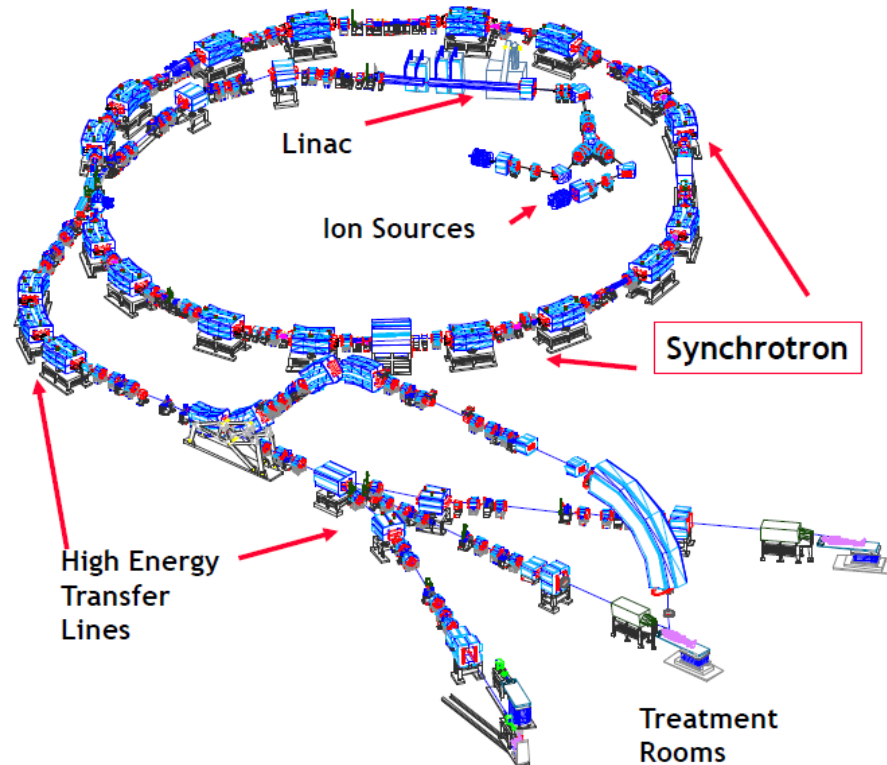
*CNAO, National Center for Oncological Hadrontherapy, Pavia, Italy*

**ENLIGHT  
ANNUAL MEETING**

CAEN, FRANCE | JULY 1-3, 2019

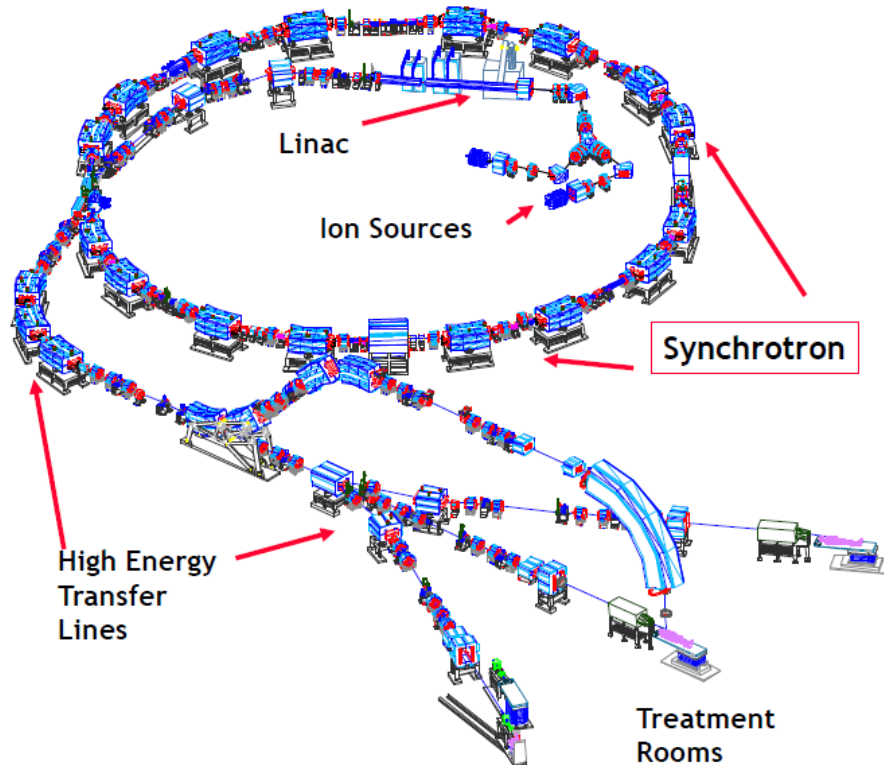


# CNAO: fixed beam lines (h&n, pelvis, abdomen, eye,...)



- ✓ 2 lateral treatment rooms → fixed horizontal beam line
- ✓ central room → horizontal and vertical fixed beam lines

# CNAO: fixed beam lines (h&n, pelvis, abdomen, eye,...)



Number of treatments with gating @ CNAO:  
*september 2014 to january 2019*

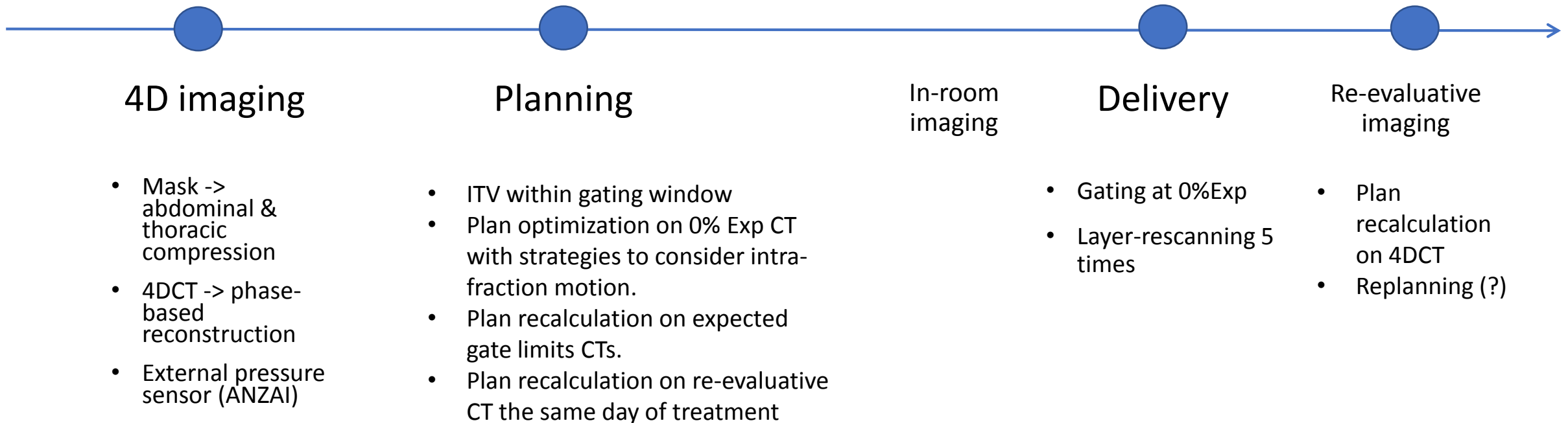
TOT	115
Upper abdomen	46
Thoracic, spine	48
Chest	21
Tumor type: pancreatic adenocarcinoma, HCC, sarcomas	

- ✓ 2 lateral treatment rooms → fixed horizontal beam line
- ✓ central room → horizontal and vertical fixed beam lines

## This talk:

- Upper abdomen moving targets.
- Pencil-beam scanning carbon ion RT plans, optimized with Siemens Syngo TPS, delivered with gating + rescanning.

## Workflow



4D imaging

Planning

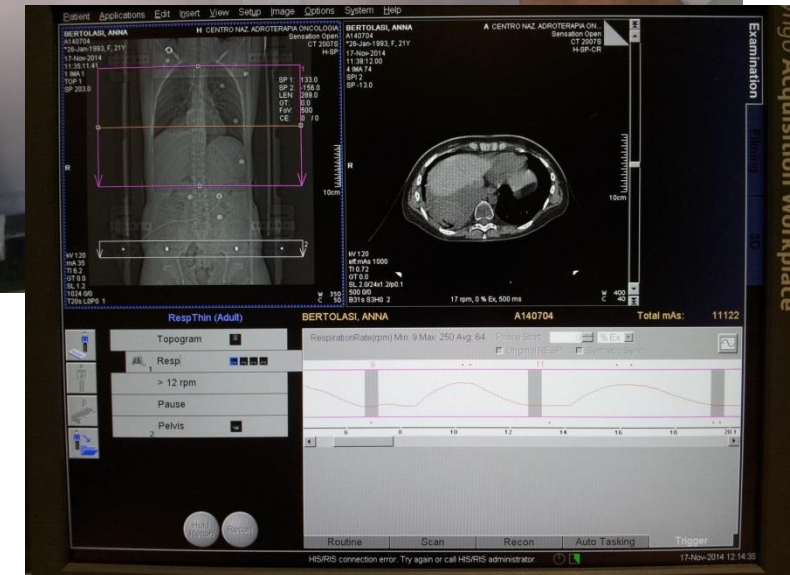
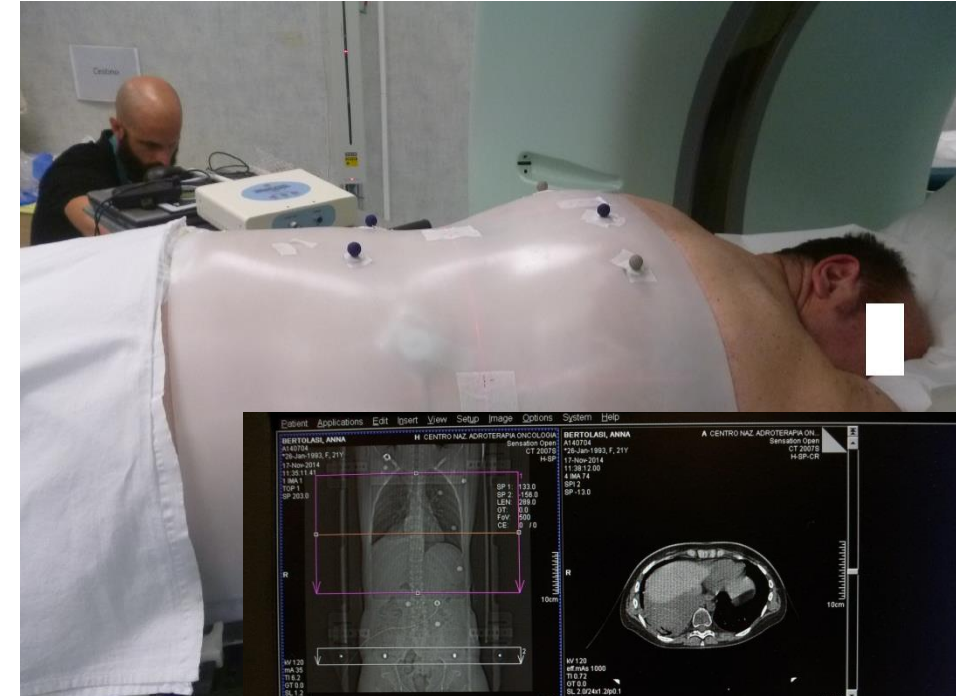
In-room imaging

Delivery

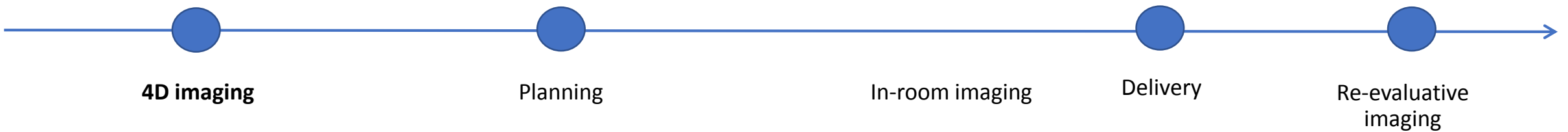
Re-evaluative  
imaging

### Characteristics:

- Mask -> abdominal & thoracic compression
  - 30%Ex:  $(0.3 \pm 0.2)$  cm, range (0.0 - 0.9)cm;
  - 30%In:  $(0.4 \pm 0.3)$  cm, range (0.0 – 1.0) cm;
- Supine and prone position
  - think at beam geometry (multiple ports)
  - consider multiple plans
- External pressure sensor (ANZAI) -> phase-based reconstruction
- 4DCT + 4DCT with MDC for organs delineation
- 5 (10) phases reconstructed (motion amplitude dependent: 0%EX,  $\pm 30\%$ ,  $\pm 50\%$ ,  $\pm 70\%$ , 90%IN )







### Characteristics:

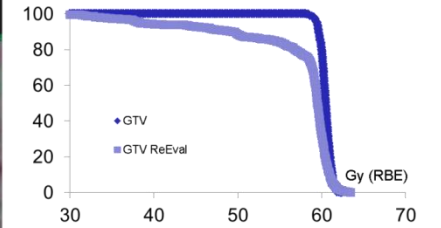
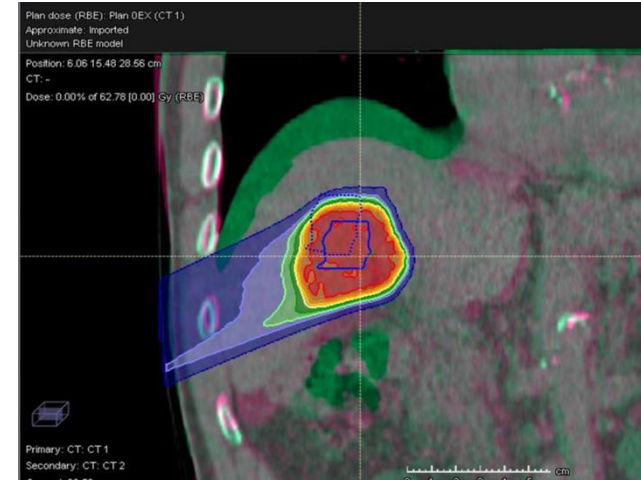
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- 5 (10) phases reconstructed (motion amplitude dependent: 0%EX,  $\pm 20\%$ ,  $\pm 30\%$ ,  $\pm 50\%$ , 90%IN )

### Limits:

- Patient setup uncertainties
- Time consuming: n contouring + n plans + n re-evaluative  
Trust deformable registration
- Trust correlation between surrogate and internal motion
- 4DCT describes an average breathing during acquisition of apprx. 1 min

## Limits: patient setup uncertainties

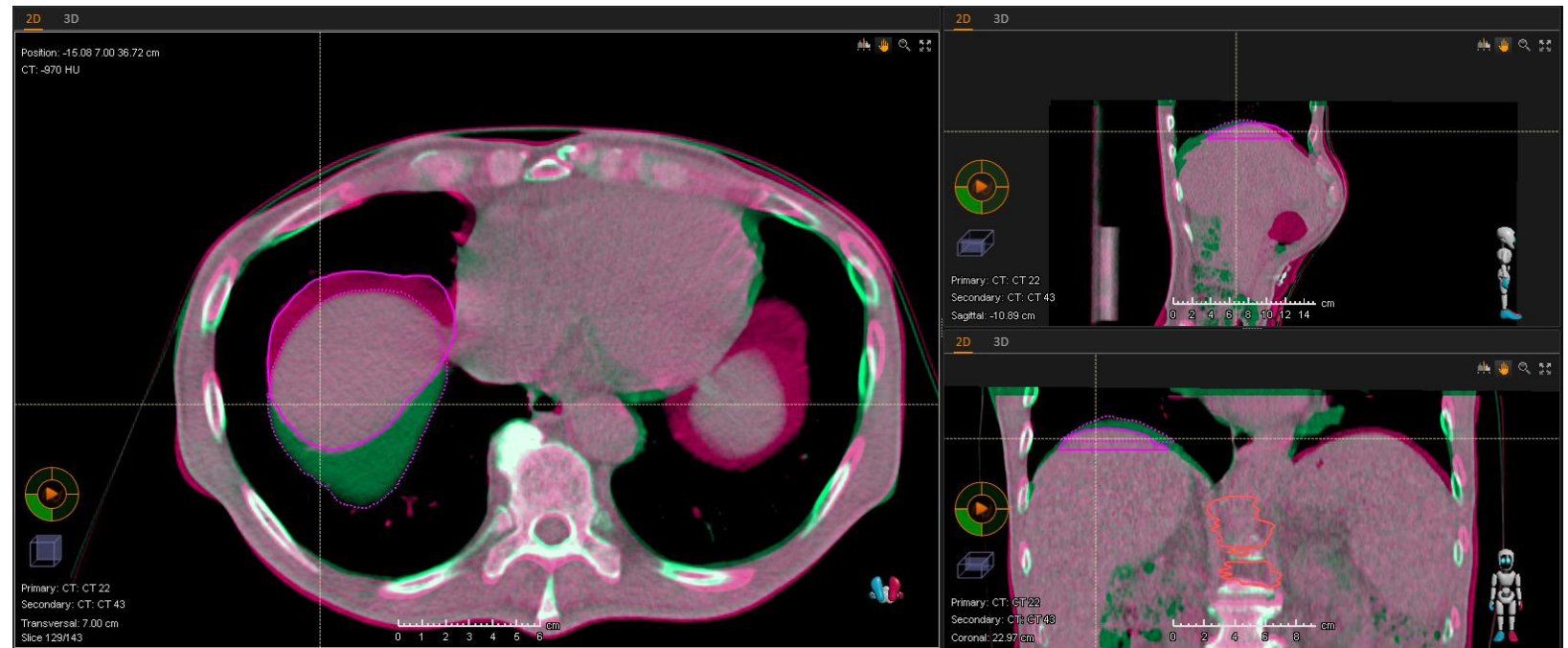
*Hepatocellular Carcinoma*  
Planning (red) vs pre-treatment (green) 0% EX CT  
NOT treated



*Pancreatic adenocarcinoma*  
Planning CT 0%EX (Dec 22, Magenta)  
Re-evaluative CT 0%ex (Jan 15, Green)

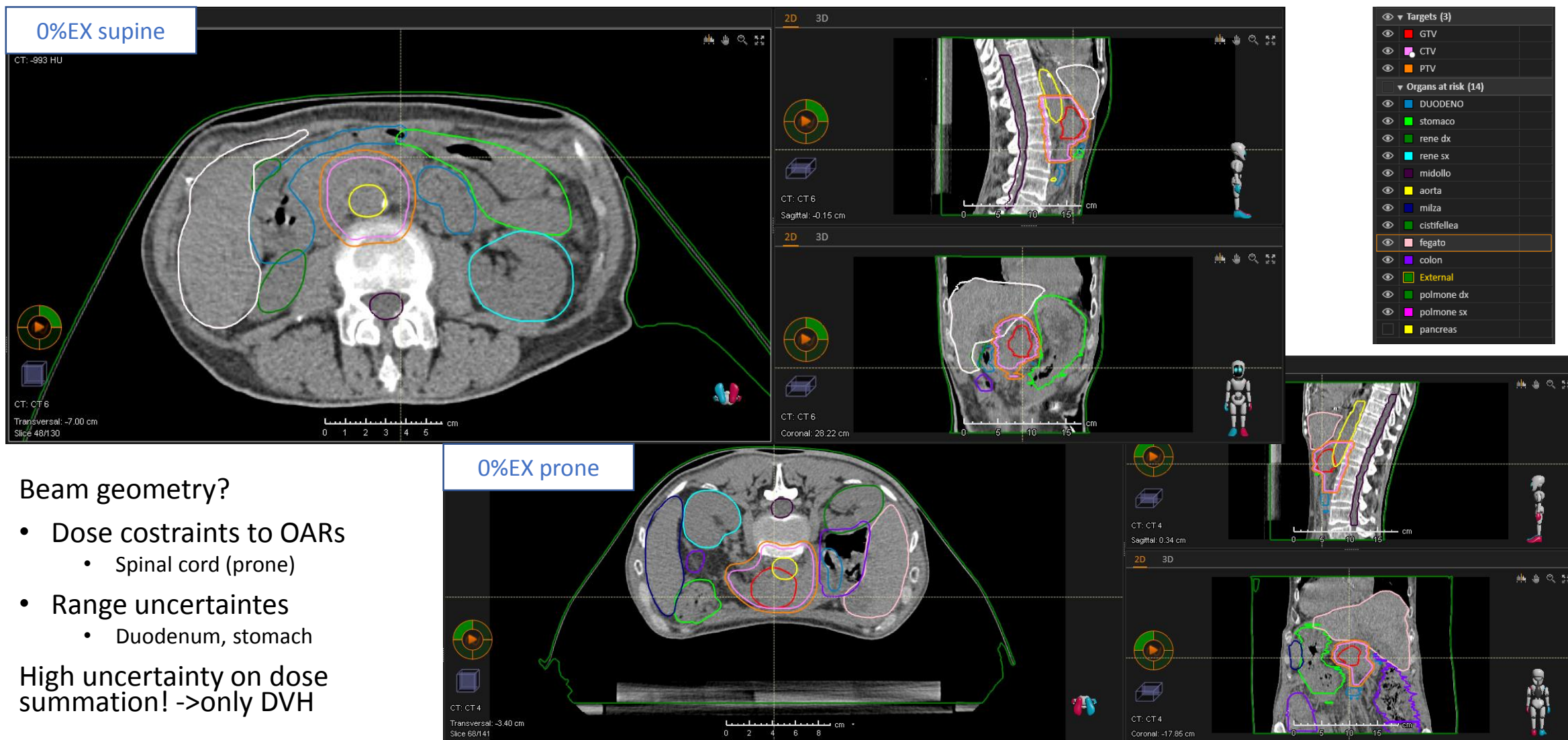
X [cm]	y[cm]	z[cm]	3D[cm]
-0.4	0.5	-1.2	1.35

*Planning CT acquired 1 day after the mask preparation?*



## Limits: multiple setups

Pancreatic adenocarcinoma, C-ion, 57.6 Gy(RBE) 12fr/4.8Gy(RBE), 9fr supine + 3 fr prone, IMPT



### Beam geometry?

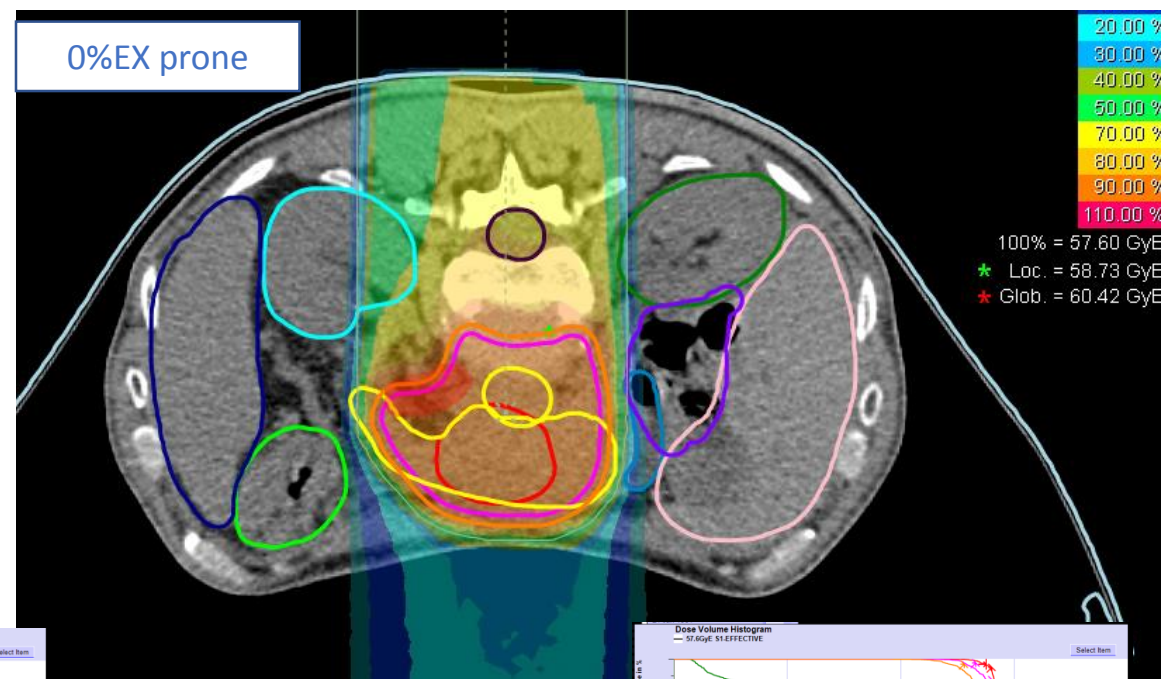
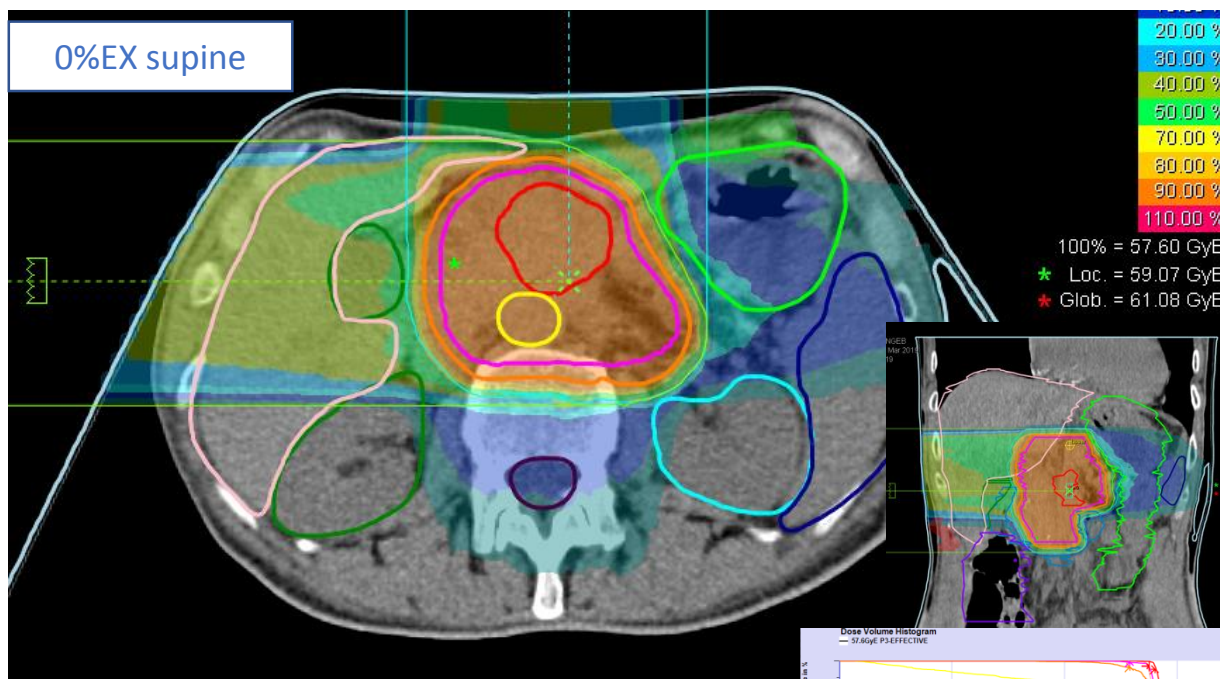
- Dose constraints to OARs
  - Spinal cord (prone)
- Range uncertainties
  - Duodenum, stomach

High uncertainty on dose summation! -> only DVH



## Multiple setups

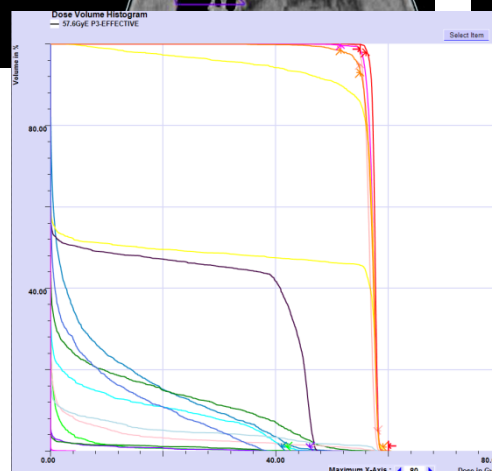
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### Beam geometry?

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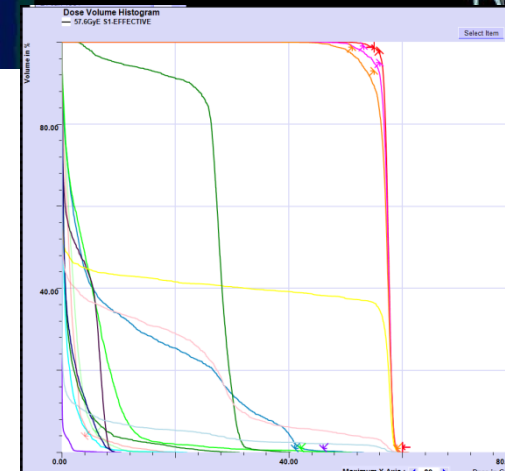
High uncertainty on dose summation! -> only DVH



9 fr

Respect on OARs dose constraints on single plans

+

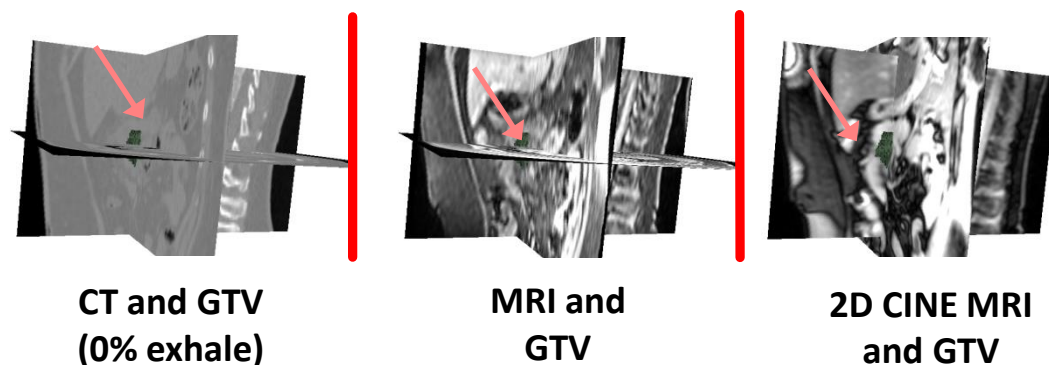


3 fr

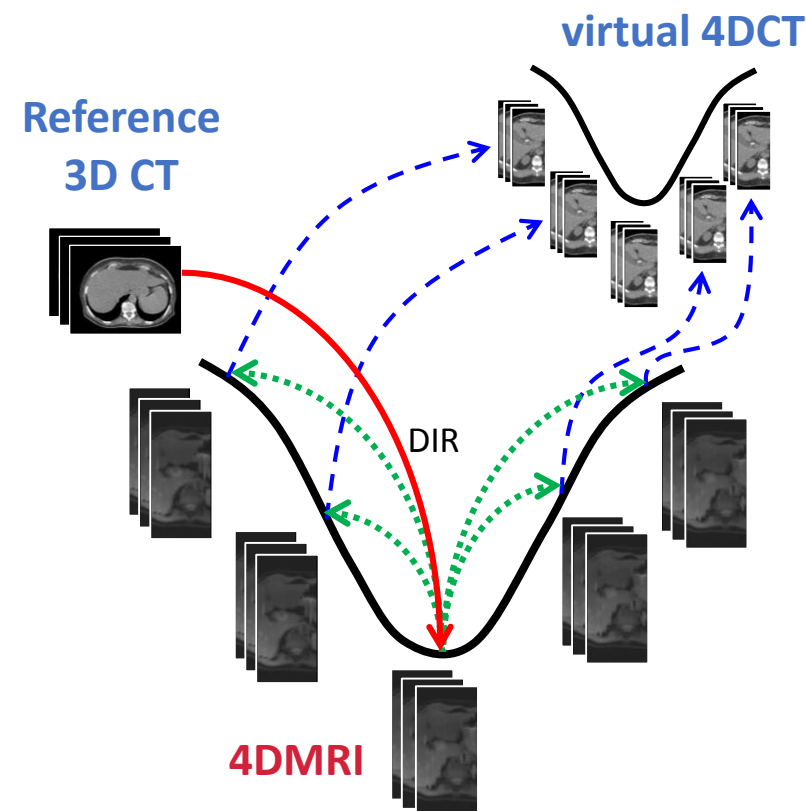
## Average respiratory cycle

Virtual 4DCT from 4D MRI

- Multi-slice acquisition of sagittal images during free-breathing (TrueFISP sequence, resolution: 1.33x1.33x5mm)
- 4DMRI retrospective sorting (Meschini et al. Phys Med 2019;58:107-13)
- 6 pancreas + 2 liver patients treated with gated carbon ion therapy
- 18 4DMRI overall (repeated acquisition during the same scan and/or after ~1 week)



[submitted manuscript]





*Purpose:* **robust** and **conformal** treatment plan

- Targets and OARs delineated on all phases: use of DIR?
- ITV within gating window (0% to  $\pm 30\%$  exp)
- Plan optimization on 0% Exp:
  - Beam geometry: avoid surfaces which moves perpendicular to beam axis;
  - Consider possible inter-fraction motion;
- Plan recalculation on different phases:
  - $\pm 30\%$  exp: estimated gating window limits
  - 100% in: worst-case scenario
- Pre-treatment recalculation (same day)



4D imaging

**Planning**

In-room imaging

Delivery

Re-evaluative  
imaging

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*Clinical case:*

Pericardial leiomyosarcoma

Carbon ion – 64Gy (RBE), 16fr/ 4 Gy(RBE)

*Datasets*

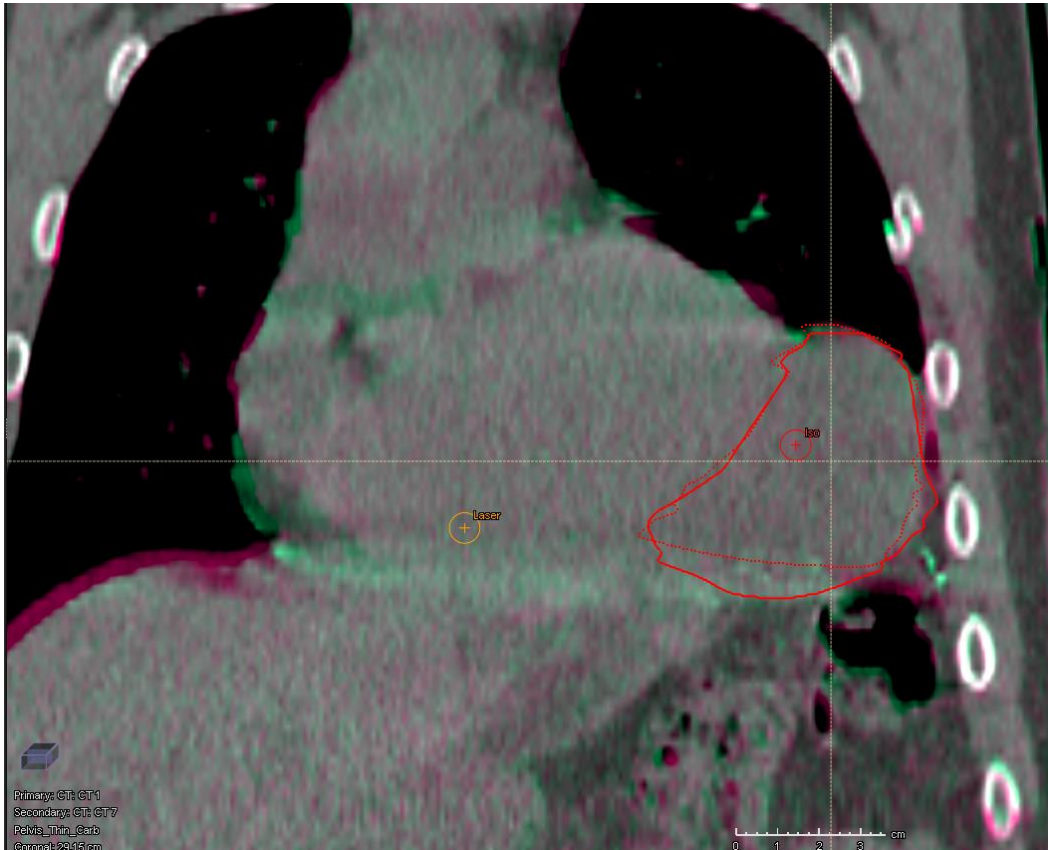
Respiratory-gated 4DCT (10 phases)

Cardiac-gated 4DMRI (external source)

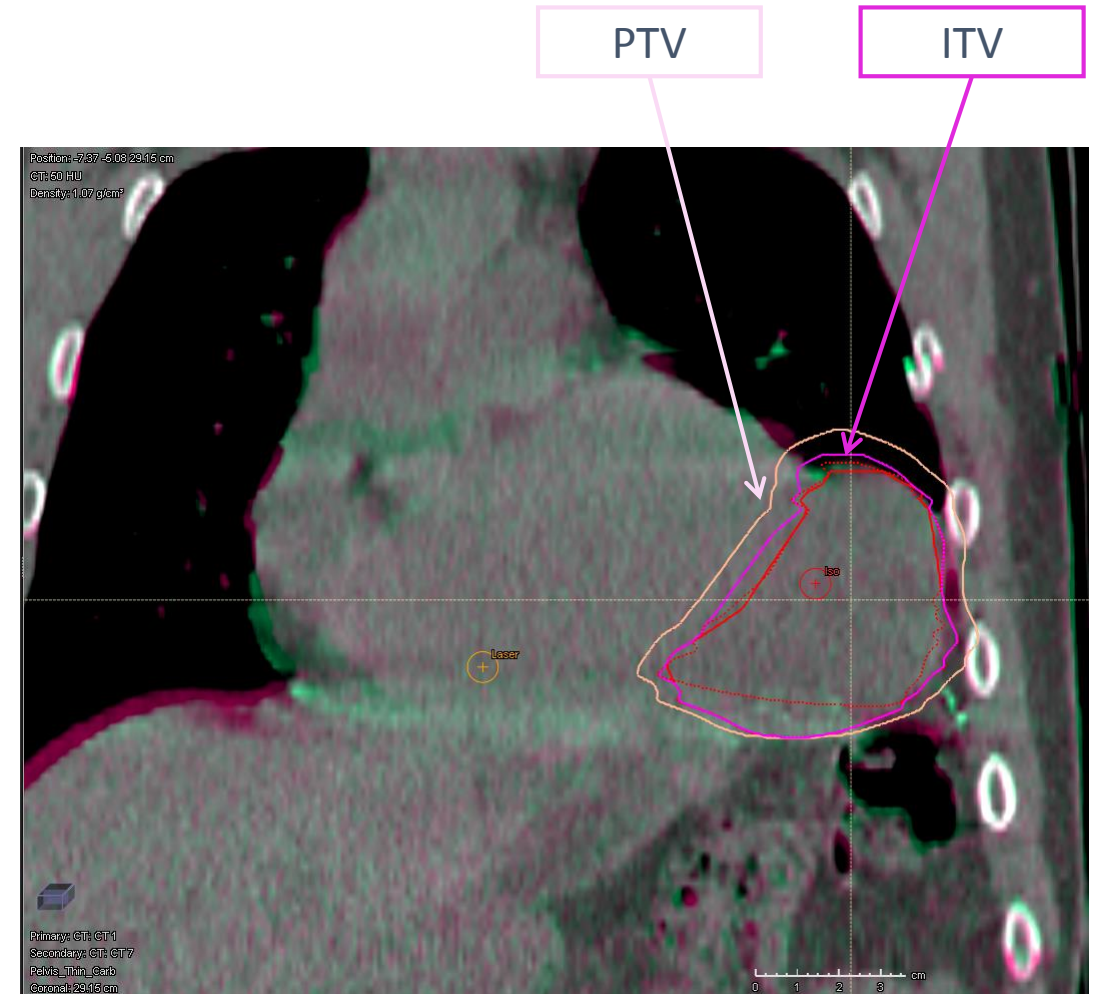


## Robust & conformal planning

Define the target



GTV 0%EX (green)  
GTV 30%EX (magenta)  
GTV 30%IN

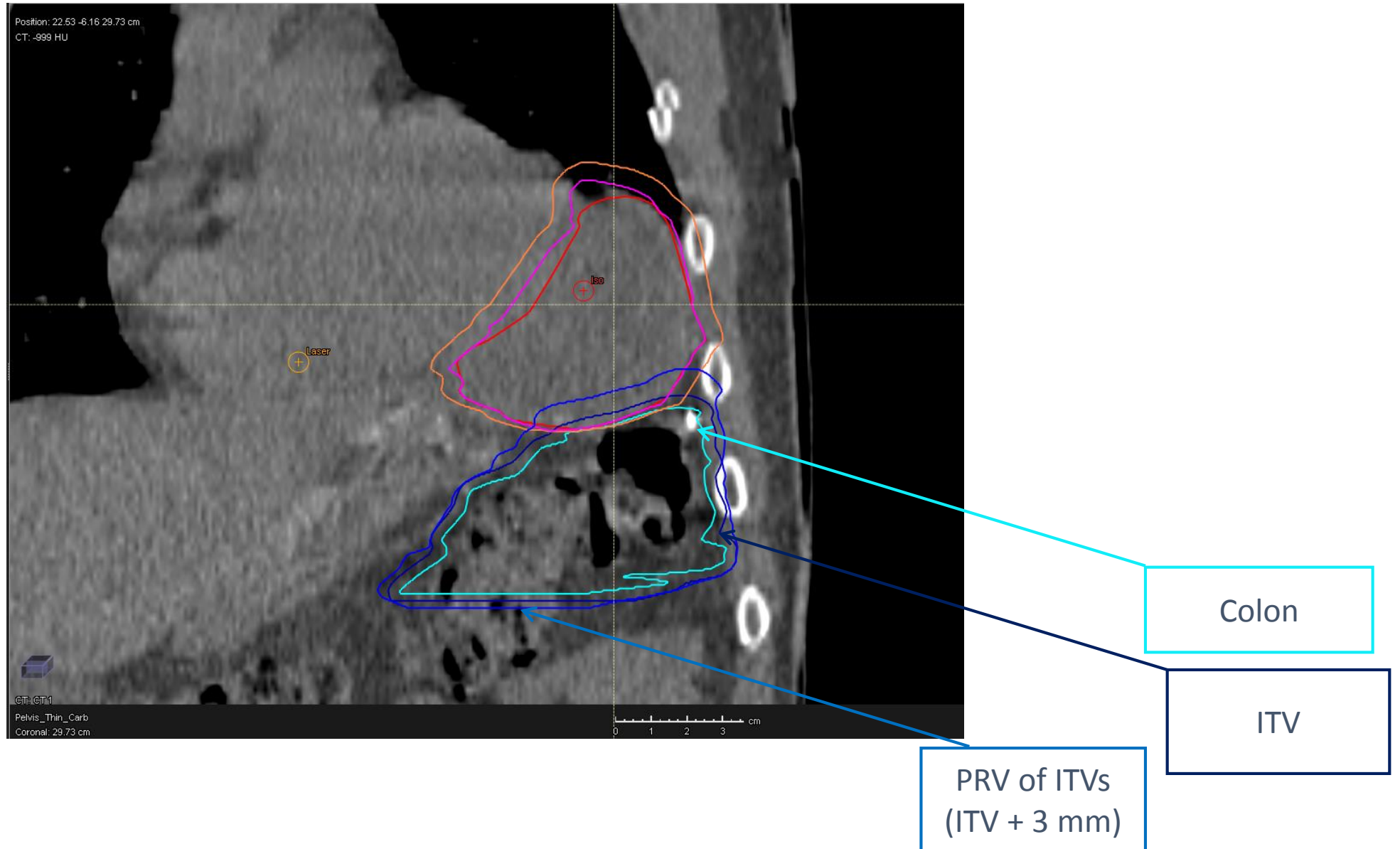


$ITV = GTV(0\%EX) + GTV(30\%IN) + GTV(30\%EX)$   
ITV manually adjusted considering 4DMRI

$PTV = ITV + 5mm$

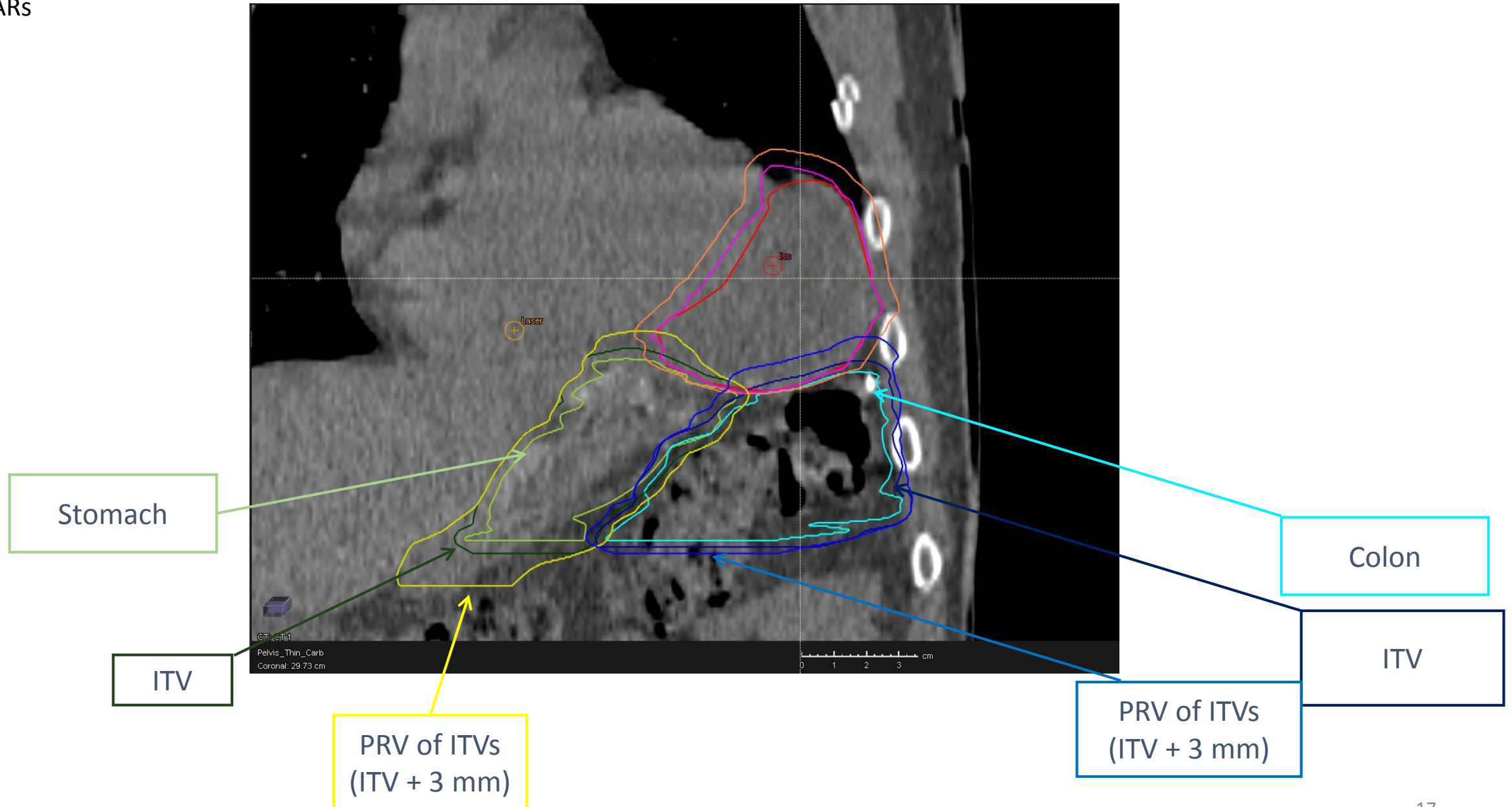
# Robust & conformal planning

OARs



# Robust & conformal planning

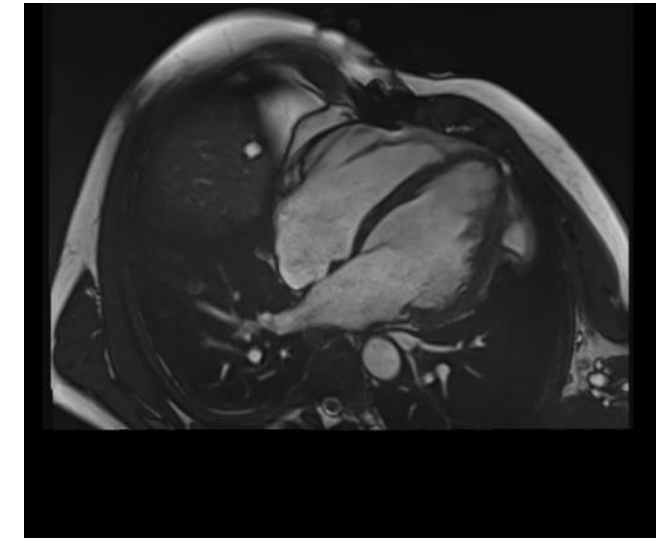
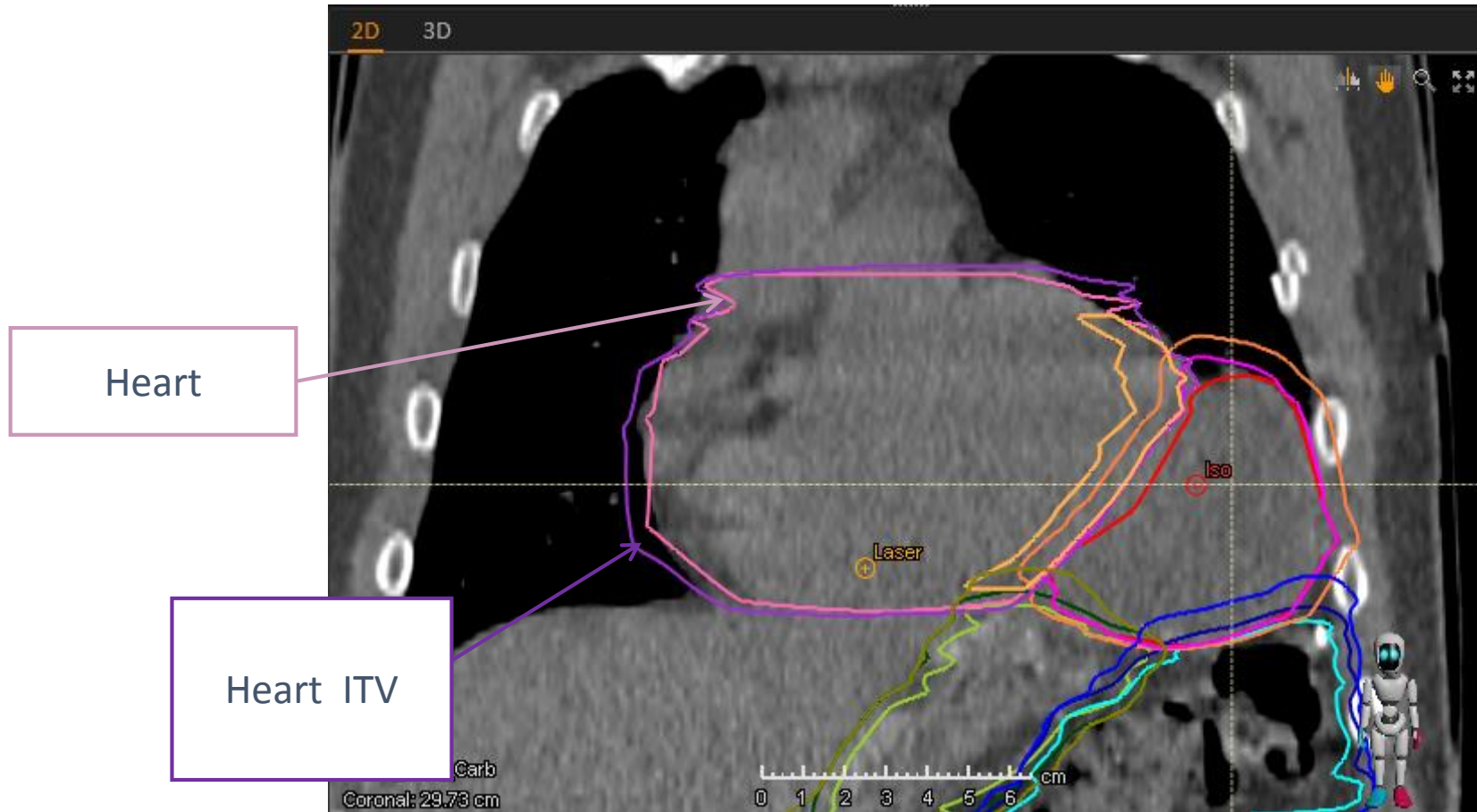
OARs





## Robust & conformal planning

OARs



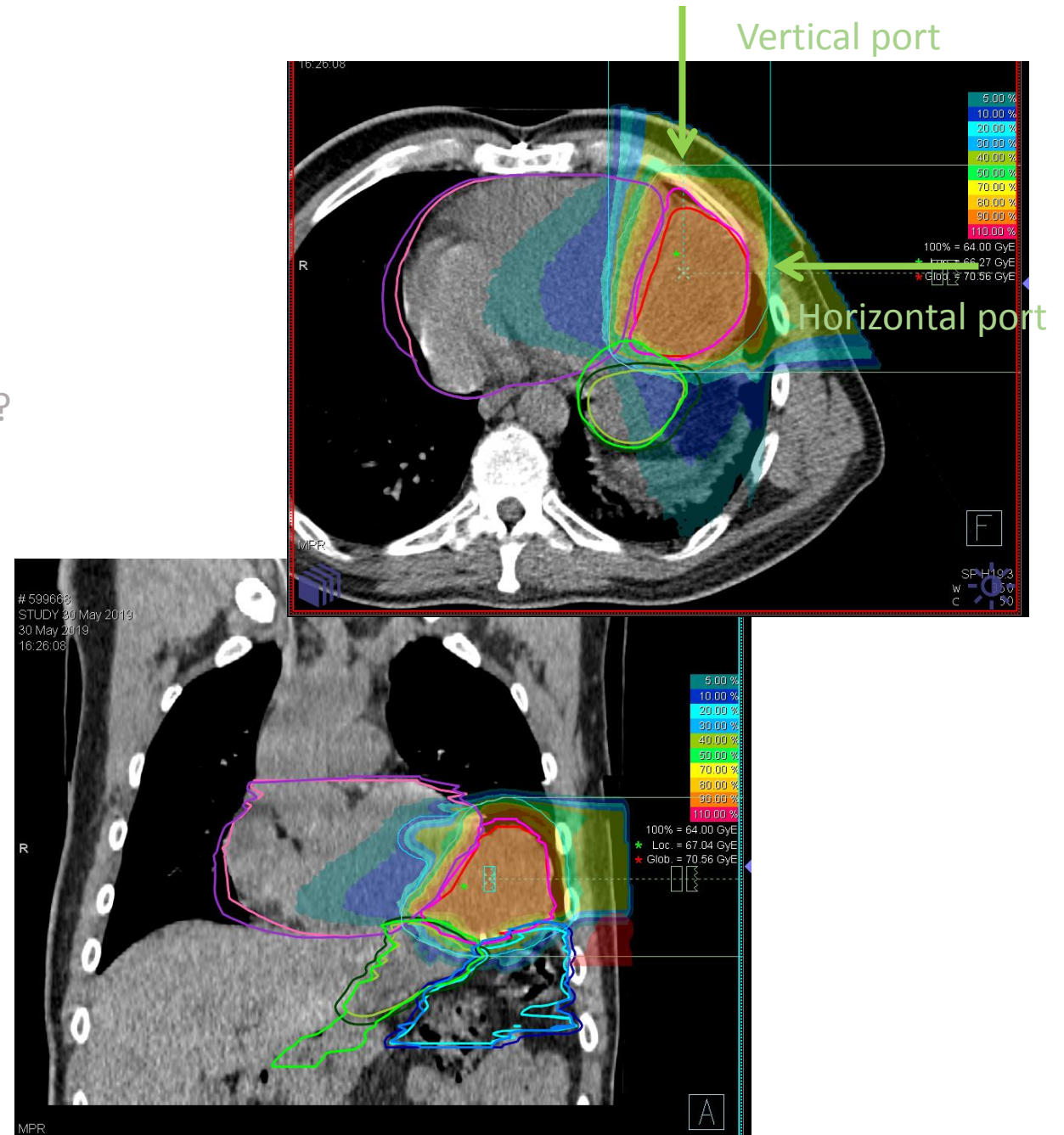
Heart: evaluate motion from external cardiac MRI -> SI and LL directions (3 mm).  
Heart ITV not used for plan optimization, only cardiac wall contour.



# Robust & conformal planning

## Plan optimization

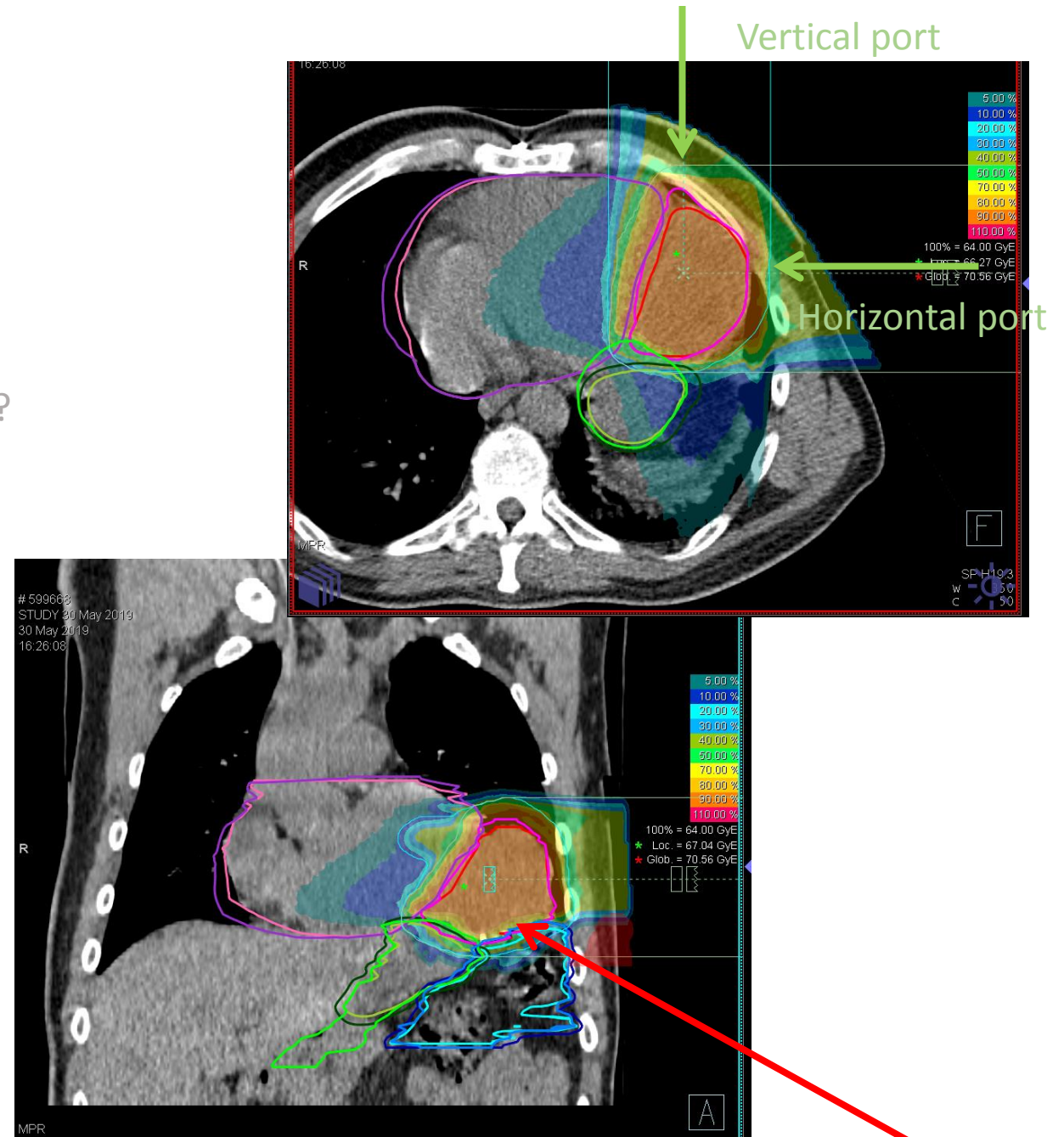
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- Pre-treatment recalculation (same day)



## Robust & conformal planning

### Plan optimization

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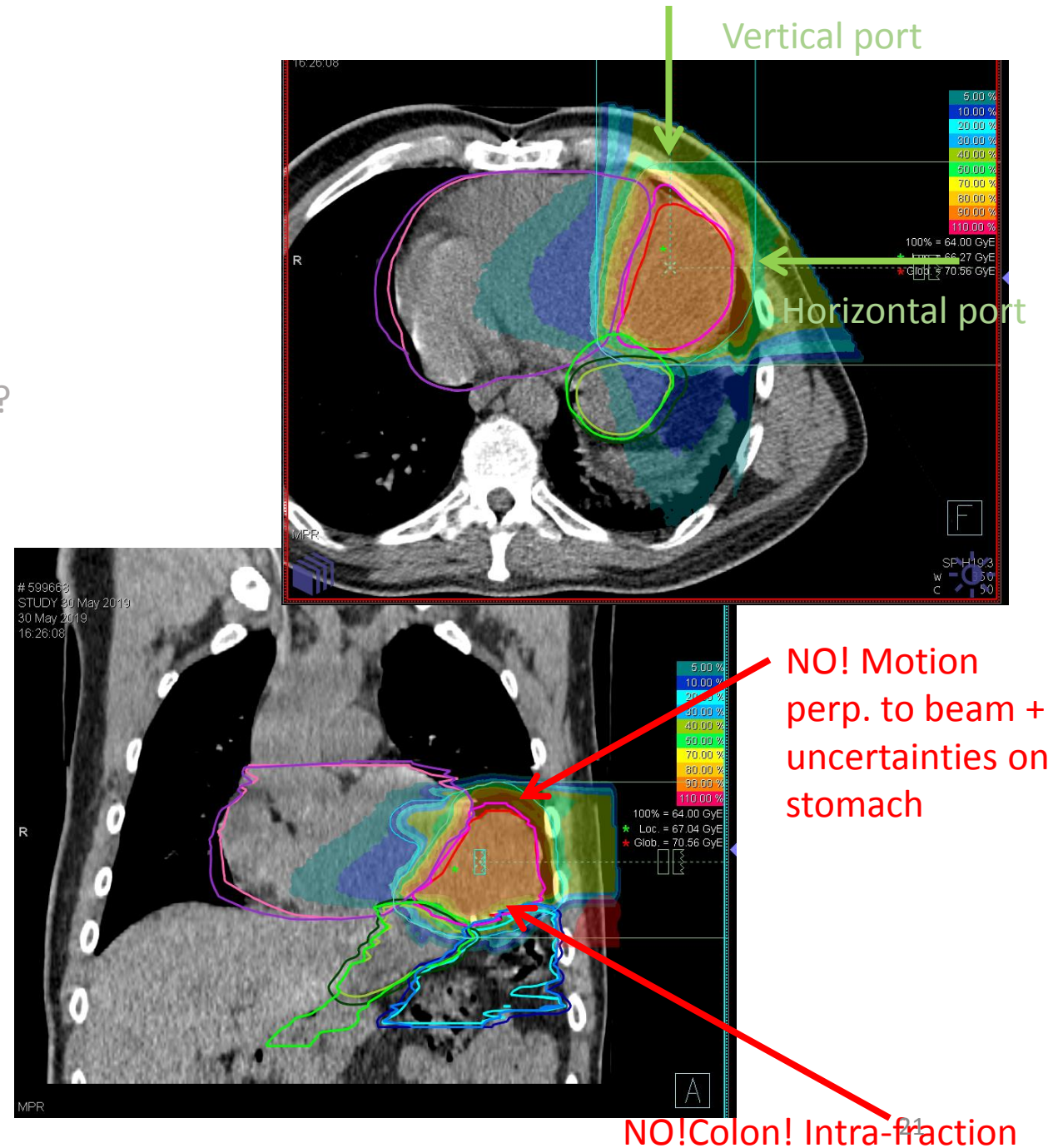


NO! Colon! Intra-fraction

## Robust & conformal planning

### Plan optimization

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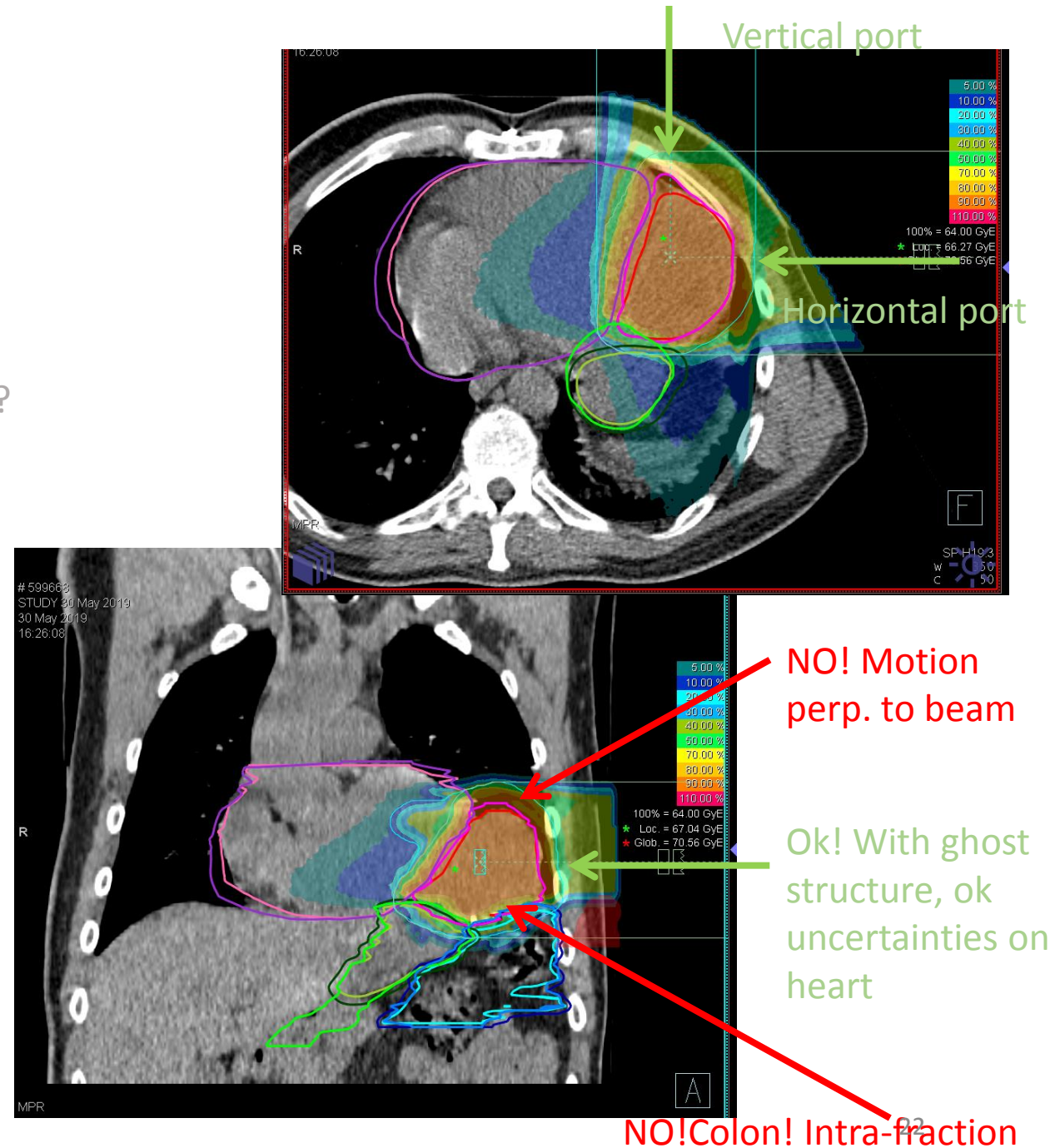




## Robust & conformal planning

### Plan optimization

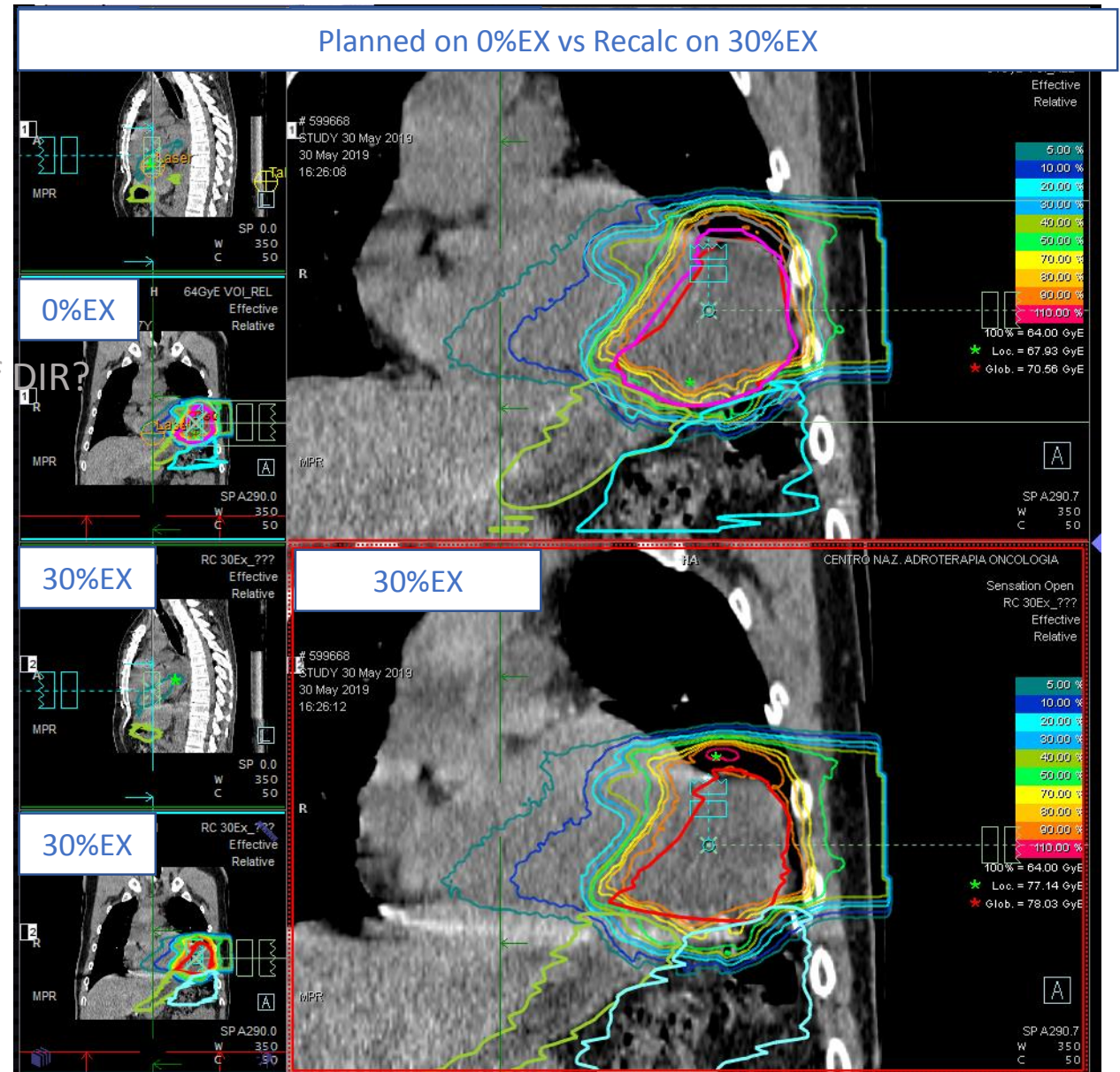
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## Plan Recalculation

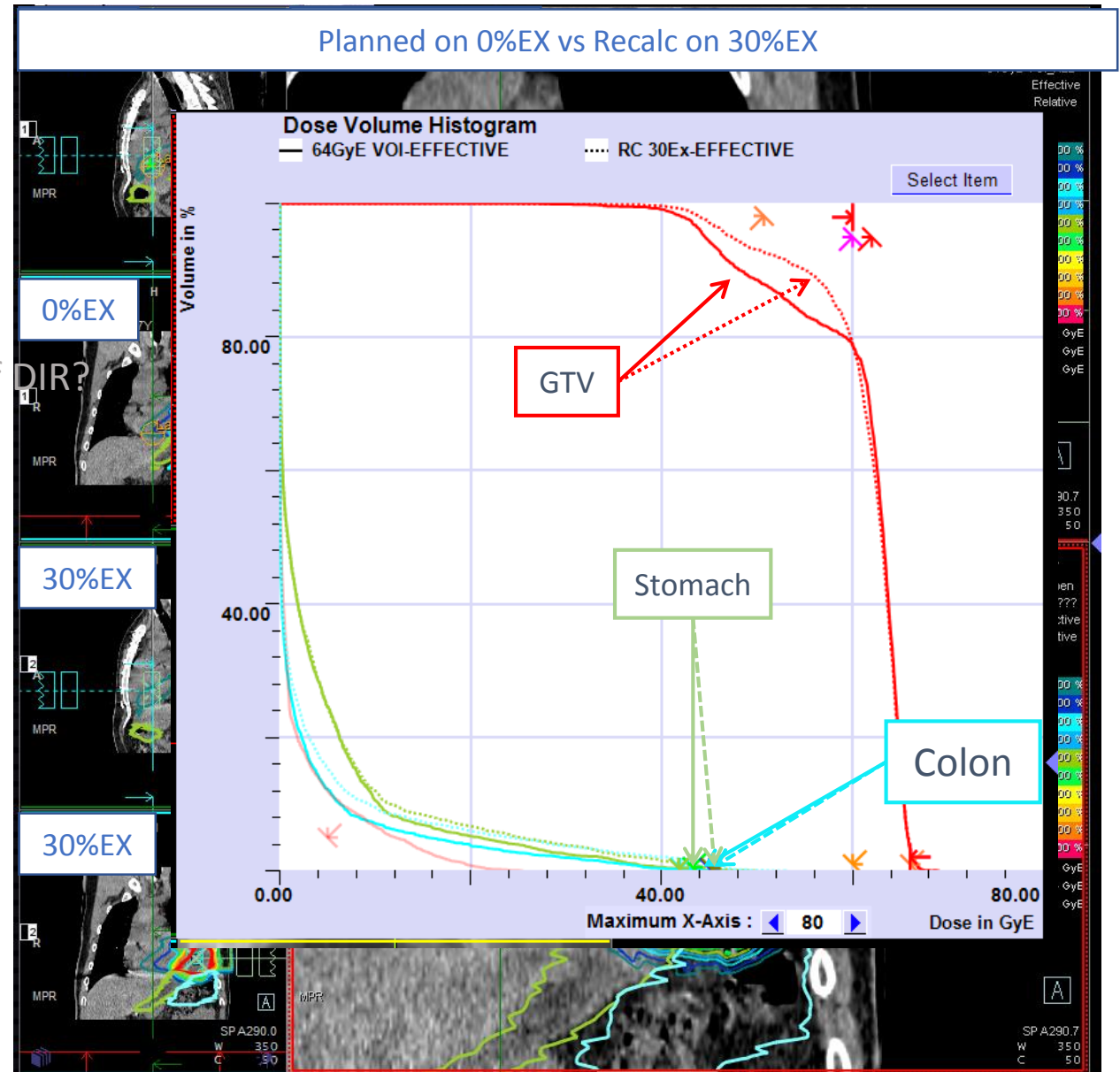
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# Robust & conformal planning

## Plan Recalculation

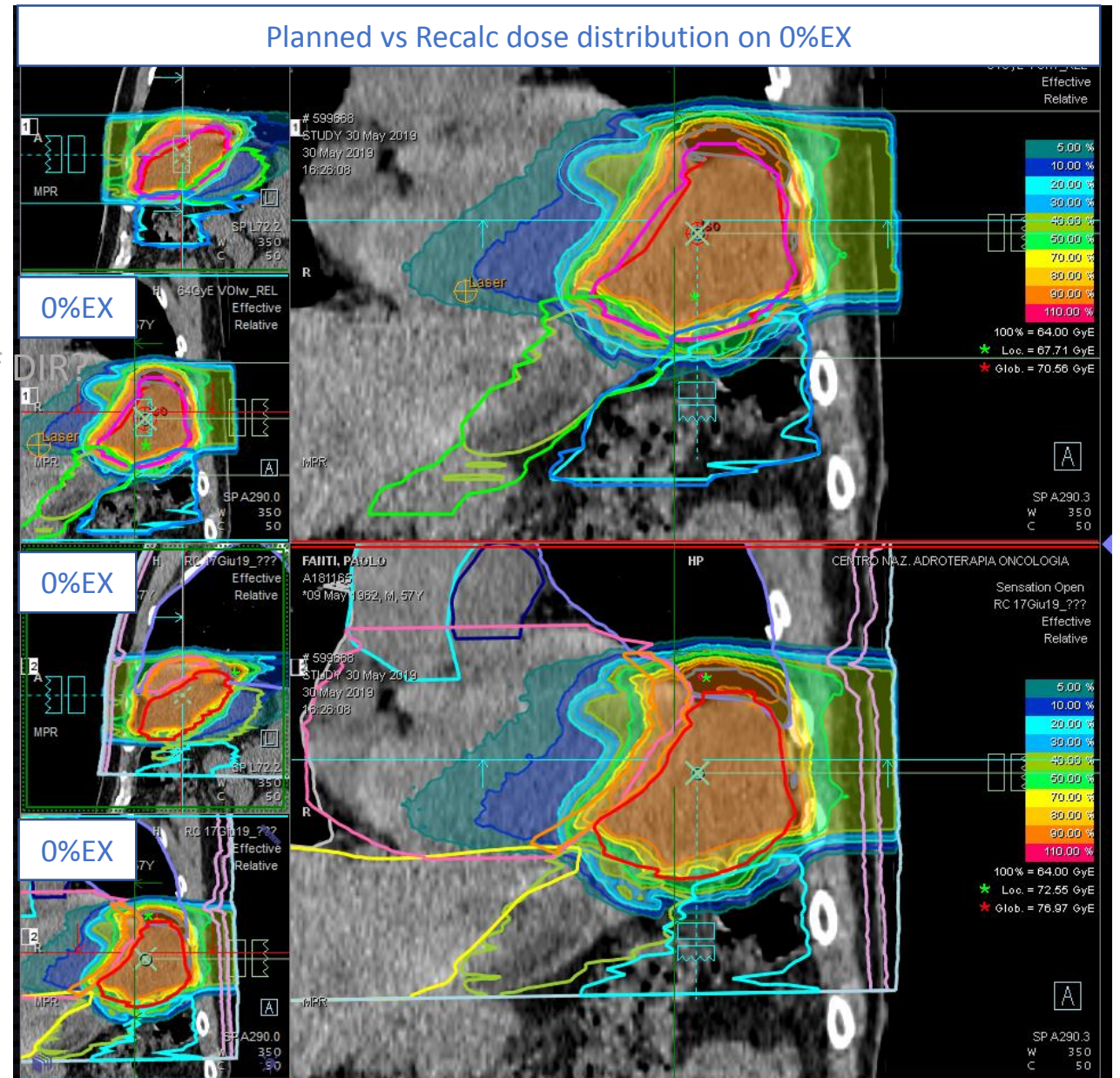
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## Plan Recalculation

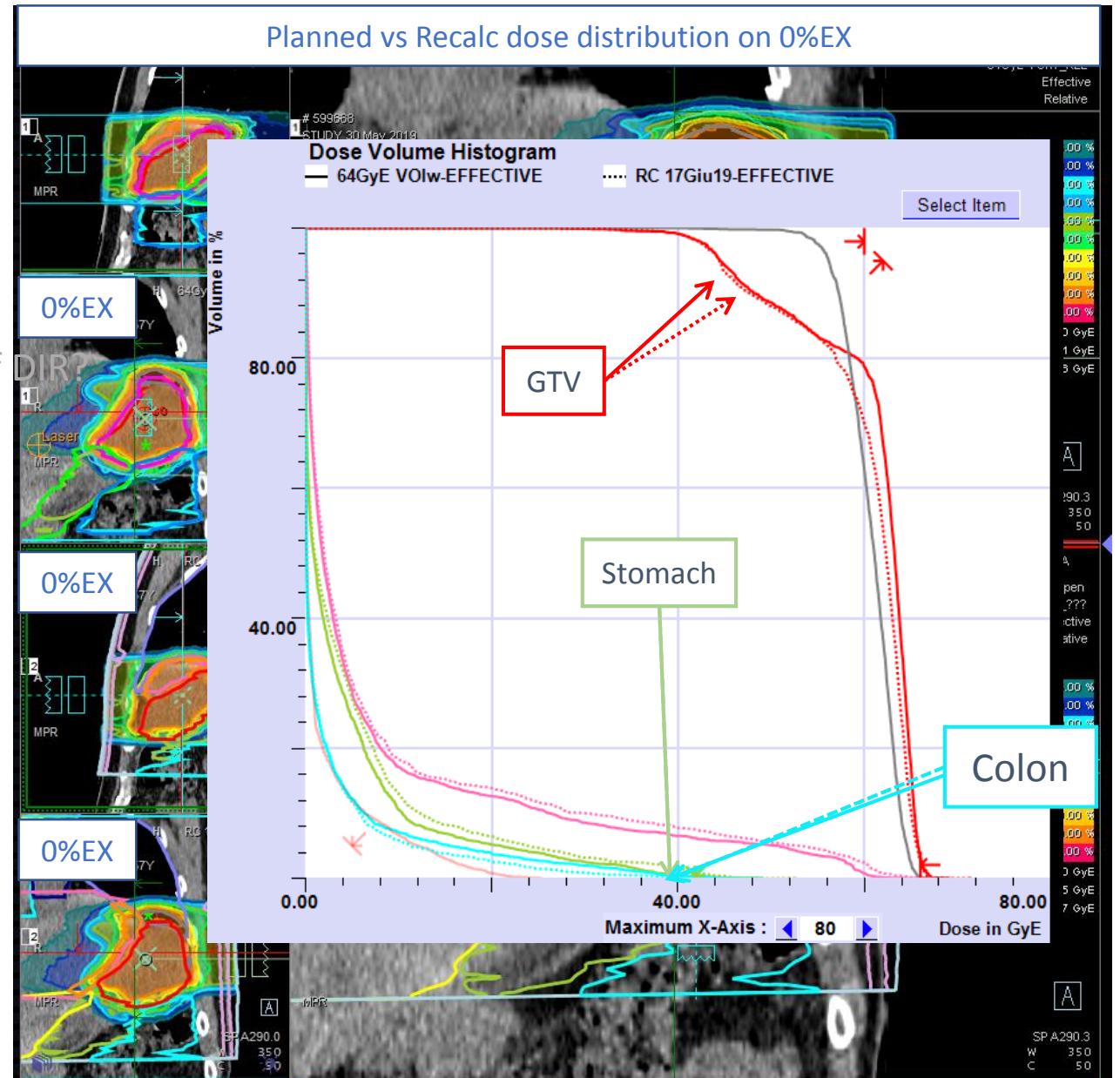
- **Pre-treatment recalculation (same day)**



# Robust & conformal planning

## Plan Recalculation

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- **Pre-treatment recalculation (same day)**



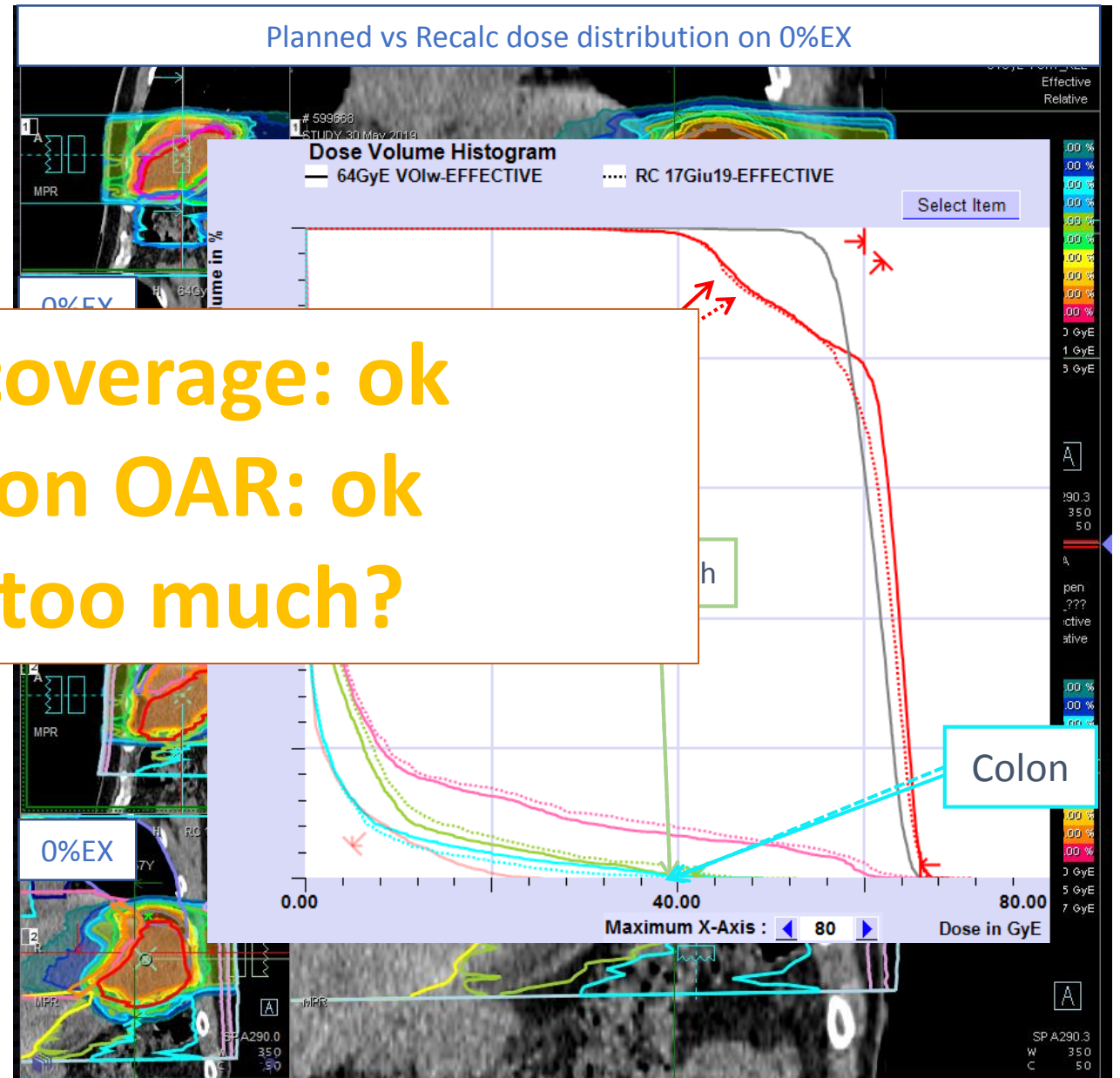


# Robust & conformal planning

## Plan Recalculation

- Targets and OARs
- ITV within gating
- Plan optimization
  - Beam geometry perpendicular
  - Consider possible
- Plan recalculation on different phases:
  - $\pm 30\%$  exp: estimated gating window limits
  - 100% in: worst-case scenario
- **Pre-treatment recalculation (same day)**

**Target coverage: ok**  
**Safety on OAR: ok**  
**Is this too much?**



4D imaging

Planning

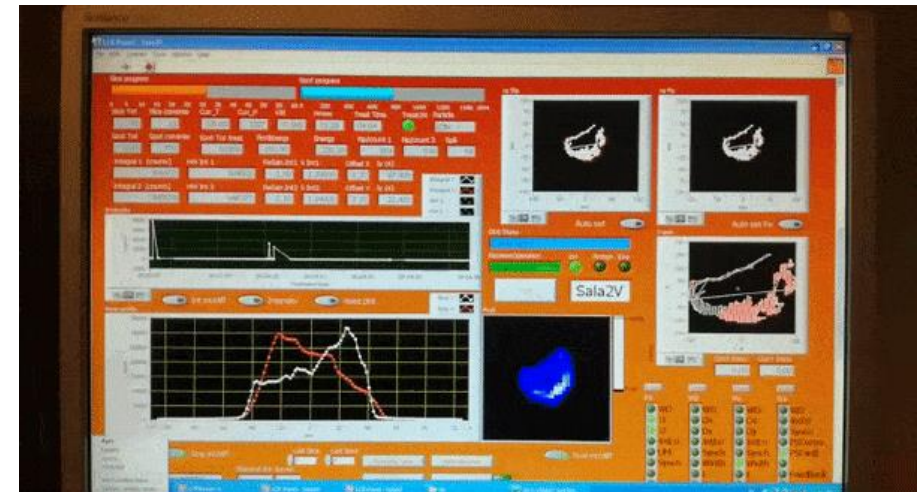
In-room imaging

**Delivery**

Re-evaluative  
imaging

## Delivery: gating + layer rescanning

- Gating at end-exhale phase.
  - Adjustable flat-top time + spill extraction synchronized with gate on
  - gate window  $\approx 1$  s
- 5-times layer rescanning
  - 5 is a compromise:
    - same number of minimum n° particles per spot as a standard plan, for each rescan
    - more rescans  $\rightarrow$  higher threshold  $\rightarrow$  less conformal dose
  - Fractionation helps vs interplay effect



Commissioning of the 4-D treatment delivery system for organ motion management in synchrotron-based scanning ion beams



Mario Ciocca<sup>a,\*</sup>, Alfredo Mirandola<sup>a</sup>, Silvia Molinelli<sup>a</sup>, Stefania Russo<sup>a</sup>, Edoardo Mastella<sup>a</sup>, Alessandro Vai<sup>a</sup>, Andrea Mairani<sup>a</sup>, Giuseppe Magro<sup>a</sup>, Andrea Pella<sup>a</sup>, Marco Donetti<sup>a</sup>, Francesca Valvo<sup>a</sup>, Piero Fossati<sup>a,b</sup>, Guido Baroni<sup>a,c</sup>



### Treatment duration: simple stats

9 patients, 27 irradiated fields

Average treat. time  $\approx$  9 min  $\rightarrow$  gating + rescanning

average QA time  $\approx$  5 min  $\rightarrow$  only rescanning

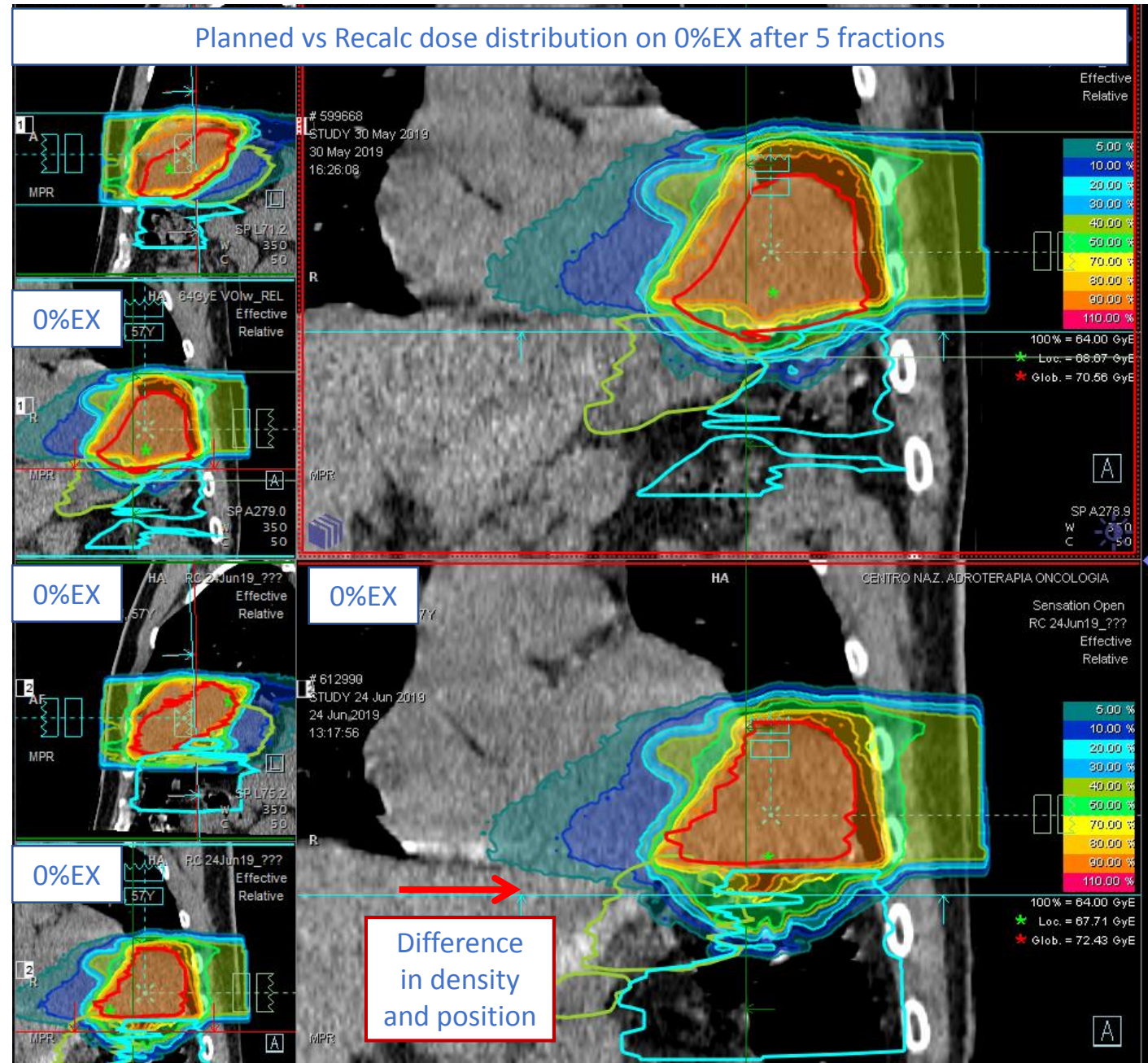
$$\frac{\text{Treat Time}}{\text{QA Time}} \approx 2.3$$



- Monitor treatment on daily imaging (CBCT, etc.) and/or re-evaluative 4DCT;



- Pericardial leiomyosarcoma  
Carbon ion – 64Gy (RBE), 16fr/ 4 Gy(RBE)

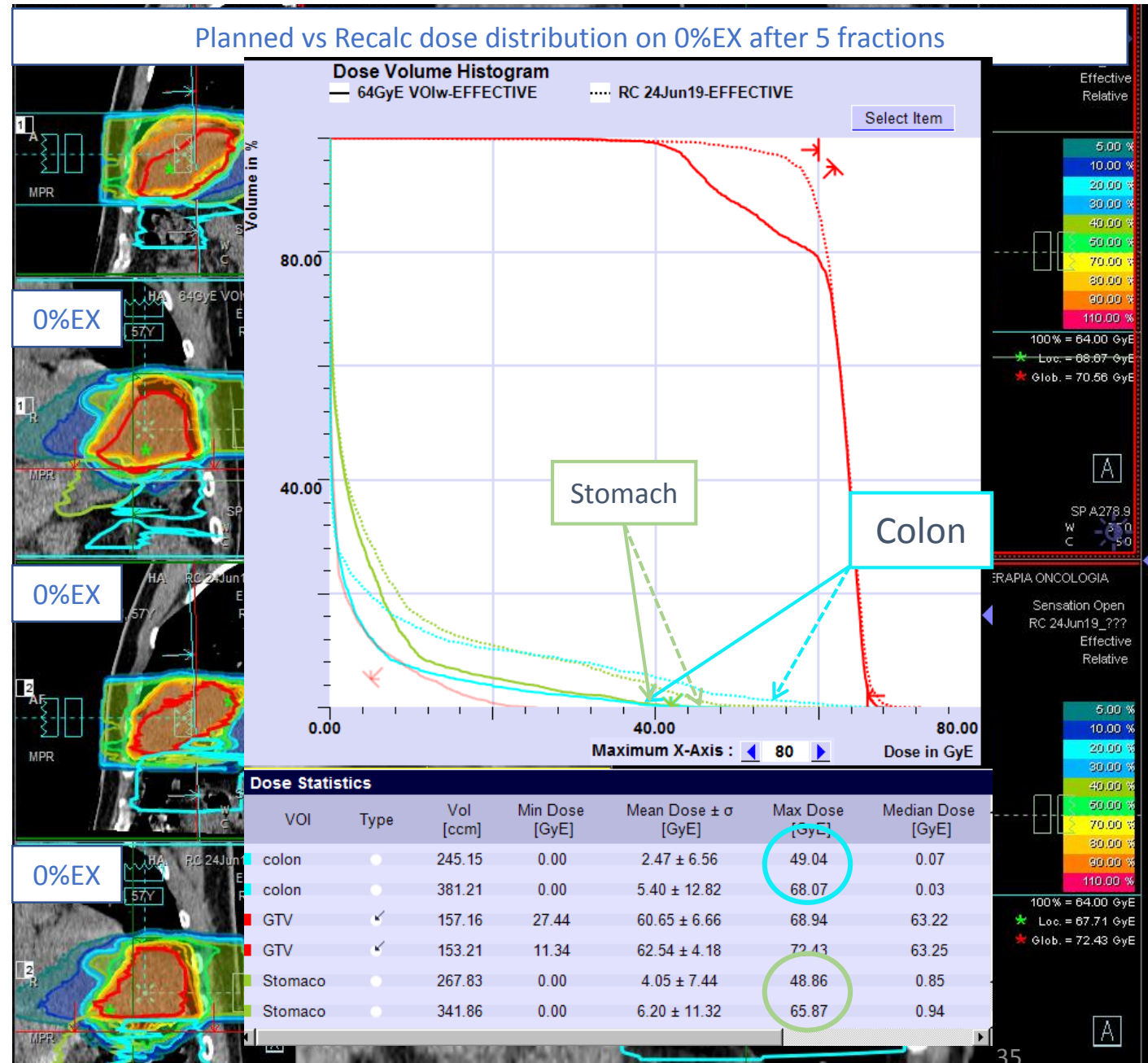


- Monitor treatment on daily imaging (CBCT, etc.) and/or re-evaluative 4DCT;

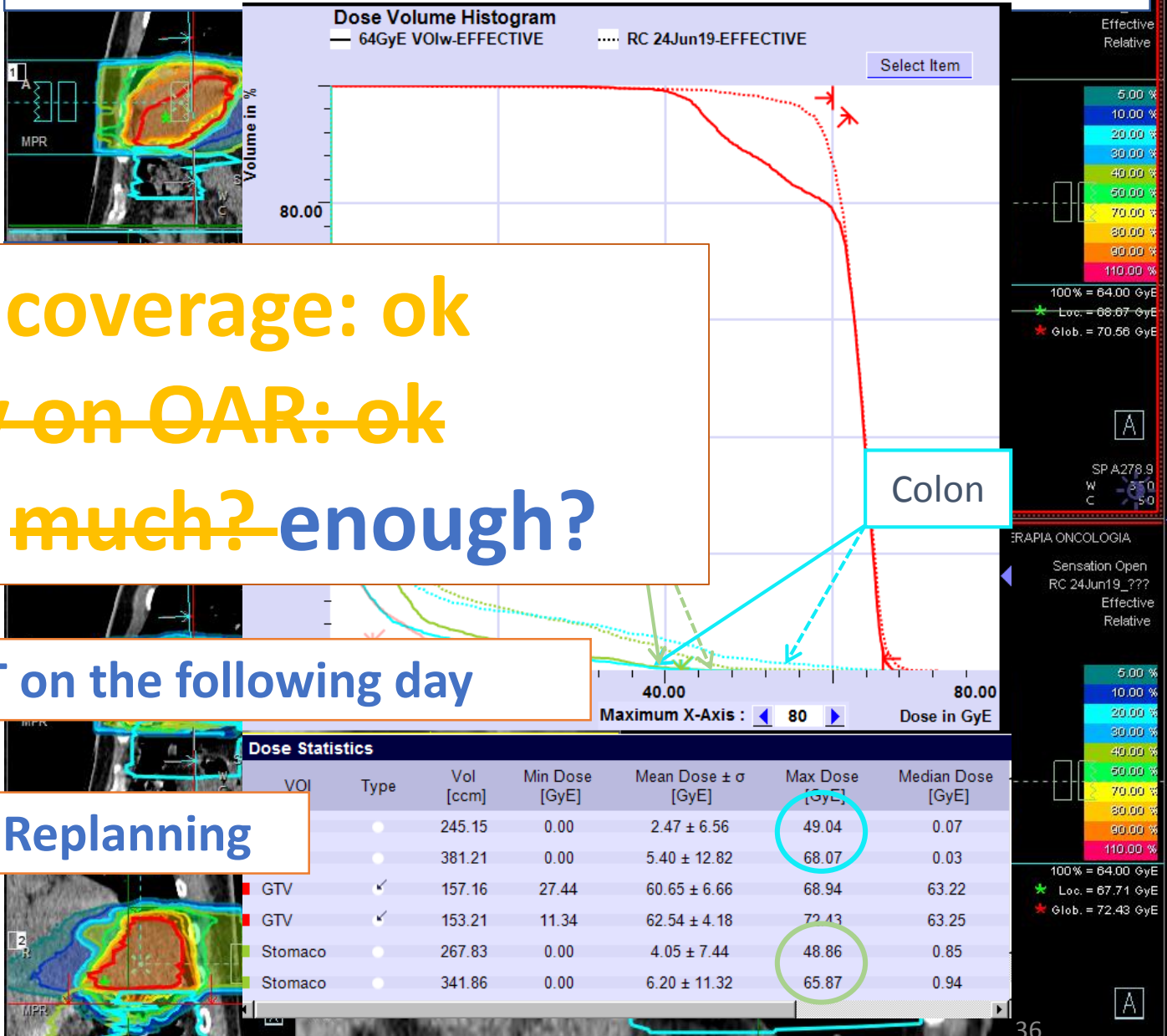
Re-evaluative 4DCT on the 2<sup>o</sup> week

Pericardial leiomyosarcoma

Carbon ion – 64Gy (RBE), 16fr/ 4 Gy(RBE)



# Planned vs Recalc dose distribution on 0%EX after 5 fractions



- Monitor treatment imaging (CBCT, etc) or re-evaluative 4D

Re-evaluative 4D

Pericardial leiomyo

Carbon ion – 64Gy (RBE), 16fr/ 4 Gy(RBE)

Target coverage: ok  
 Safety on OAR: ok  
 Is this too much? enough?

New 4DCT on the following day

Replanning



# Under development (1): robust planning

Commercial TPS with tools for **robust planning**:

- with protons (Raystation, Eclipse...)
- with carbon ions (Raystation v.8b)

The TPS simulates scenarios of setup and range uncertainties, or organ motion as represented by a 4D set, and optimize a robust plan.



# Under development (1): robust planning

Commercial TPS with tools for **robust planning**:

- with protons (Raystation, Eclipse...)
- with carbon ions (Raystation v.8b)

The TPS simulates scenarios of setup and range uncertainties, or organ motion as represented by a 4D set, and optimize a robust plan.

## Minimax optimization for handling range and setup uncertainties in proton therapy

Albin Fredriksson<sup>a)</sup>

*Department of Mathematics, Optimization and Systems Theory, Royal Institute of Technology (KTH), SE-100 44 Stockholm, Sweden and RaySearch Laboratories, Sveavägen 75, SE-111 34 Stockholm, Sweden*

Anders Forsgren

*Department of Mathematics, SE-100 44 Stockholm, Sweden*

Björn Hårdemark

*RaySearch Laboratories, Sve*

**Methods:** Dose contributions for a number of range and setup errors are calculated and a minimax optimization is performed. The minimax optimization aims at minimizing the penalty of the worst case scenario. Any optimization function from conventional treatment planning can be utilized by the method. By considering only scenarios that are physically realizable, the unnecessary conser-

# Under development (1): robust planning

**Robustness Setting**

Set robustness parameters (density, setup errors + 4d images) for optimization

Patient position uncertainty

☒ Use isotropic uncertainty

Superior [cm] 0.30  
Right [cm] 0.30  
Posterior [cm] 0.30  
Anterior [cm] 0.30  
Left [cm] 0.30  
Inferior [cm] 0.30

Position uncertainty setting

☒ Universal  
☐ Independent beams  
☐ Independent isocenters

The position uncertainties can be applied universally for all beams, or independently to each beam or isocenter in specific directions. The number of scenarios increases exponentially with the number of independent beams (isocenters), leading to longer computation times.

Range uncertainty

Range uncertainty [%]: 3.00

The range uncertainty is modeled by scaling the mass density of the patient. The range uncertainty is universal for all beams.

Accurate scenario doses

☒ Compute accurate scenario doses

Select this option to compute accurate spot doses for each robustness scenario with the chosen optimization dose engine. Otherwise, an approximate dose engine will be used for all scenarios except the nominal (unshifted) scenario on each selected image set.

Scenarios to compute: 105

Image sets

- ☐ 4D
- ☒ 4DCT\_11aprile
  - ☒ CT: CT 0EX [11 Apr 2019, 16:10:29 (hr:min:sec)]
  - ☒ CT: CT 10IN [11 Apr 2019, 16:10:30 (hr:min:sec)]
  - ☒ CT: CT 20IN [11 Apr 2019, 16:10:30 (hr:min:sec)]
  - ☐ CT: CT 30IN [11 Apr 2019, 16:10:30 (hr:min:sec)]
  - ☐ CT: CT 50 IN [11 Apr 2019, 16:10:30 (hr:min:sec)]
  - ☐ CT: CT 90IN [11 Apr 2019, 16:10:30 (hr:min:sec)]
  - ☐ CT: CT 100 IN [11 Apr 2019, 16:10:31 (hr:min:sec)]
  - ☐ CT: CT 50 EX [11 Apr 2019, 16:10:28 (hr:min:sec)]
  - ☐ CT: CT 30EX [11 Apr 2019, 16:10:28 (hr:min:sec)]
  - ☒ CT: CT 20EX [11 Apr 2019, 16:10:28 (hr:min:sec)]
  - ☒ CT: CT 10EX [11 Apr 2019, 16:10:28 (hr:min:sec)]
- ☐ ReEval\_26apr
  - ☐ CT: CT 1 [09 Apr 2019, 12:45:35 (hr:min:sec)]
  - ☐ CT: CT 2 [09 Apr 2019, 12:45:35 (hr:min:sec)]
  - ☐ CT: CT 3 [09 Apr 2019, 13:23:53 (hr:min:sec)]
  - ☐ CT: CT 4 [09 Apr 2019, 13:23:53 (hr:min:sec)]
  - ☐ CT: CT LOCKBAR [11 Apr 2019, 16:12:44 (hr:min:sec)]
  - ☐ CT: CT 13 [26 Apr 2019, 09:51:26 (hr:min:sec)]

Select all Select none

OK Cancel

**Evaluate robustness on calculated dose distribution**

Select Beam Set:

Dose properties

Density perturbation [%]: 1.0

Isocenter shift X [cm]: 0.10

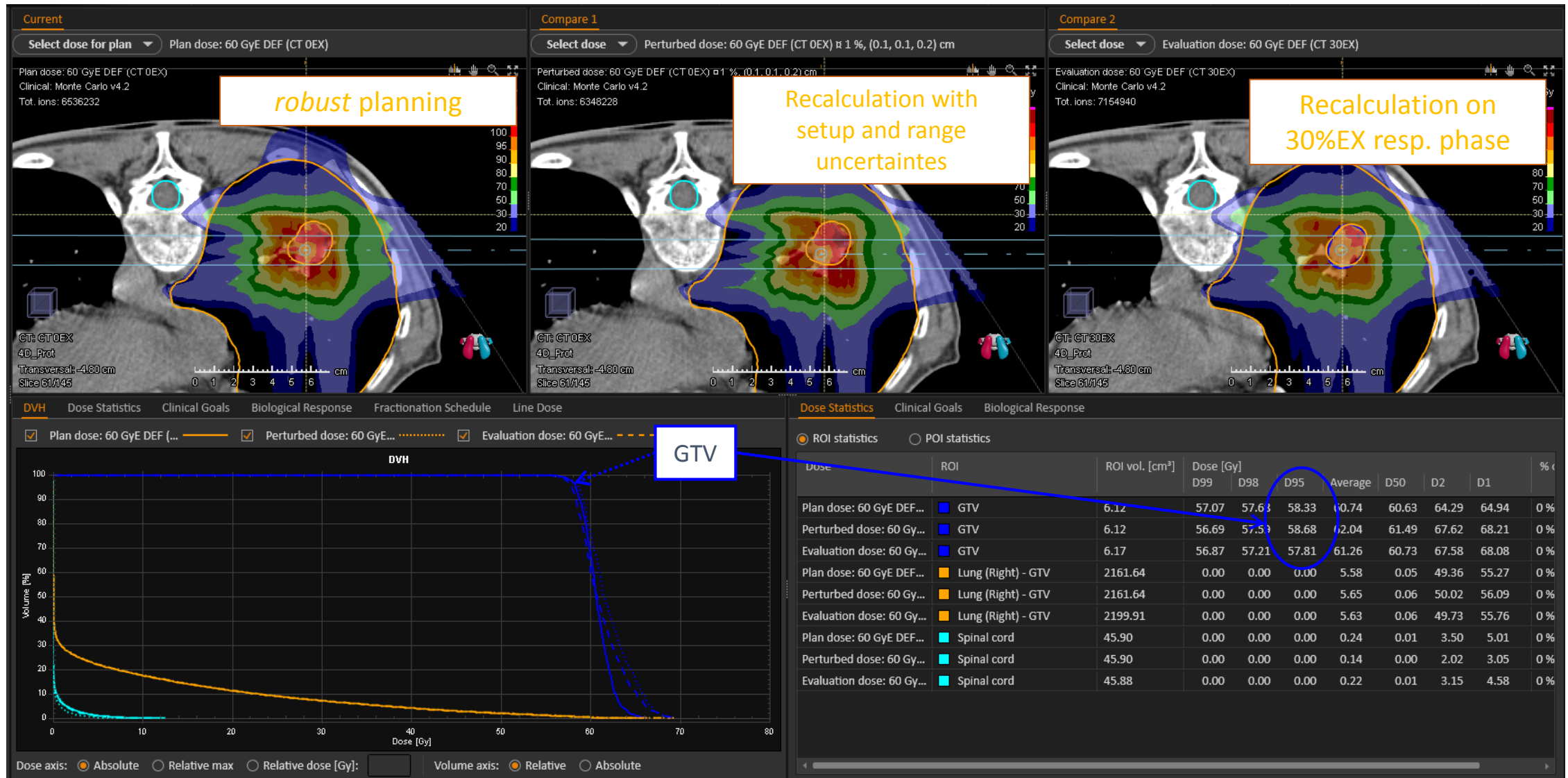
Isocenter shift Y [cm]: 0.10

Isocenter shift Z [cm]: 0.2

OK Cancel

# Robust planning

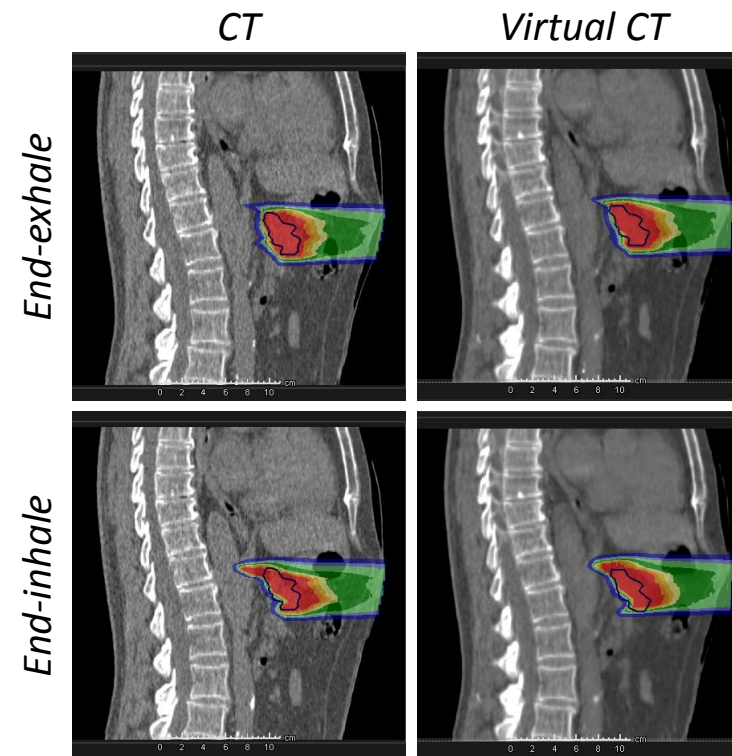
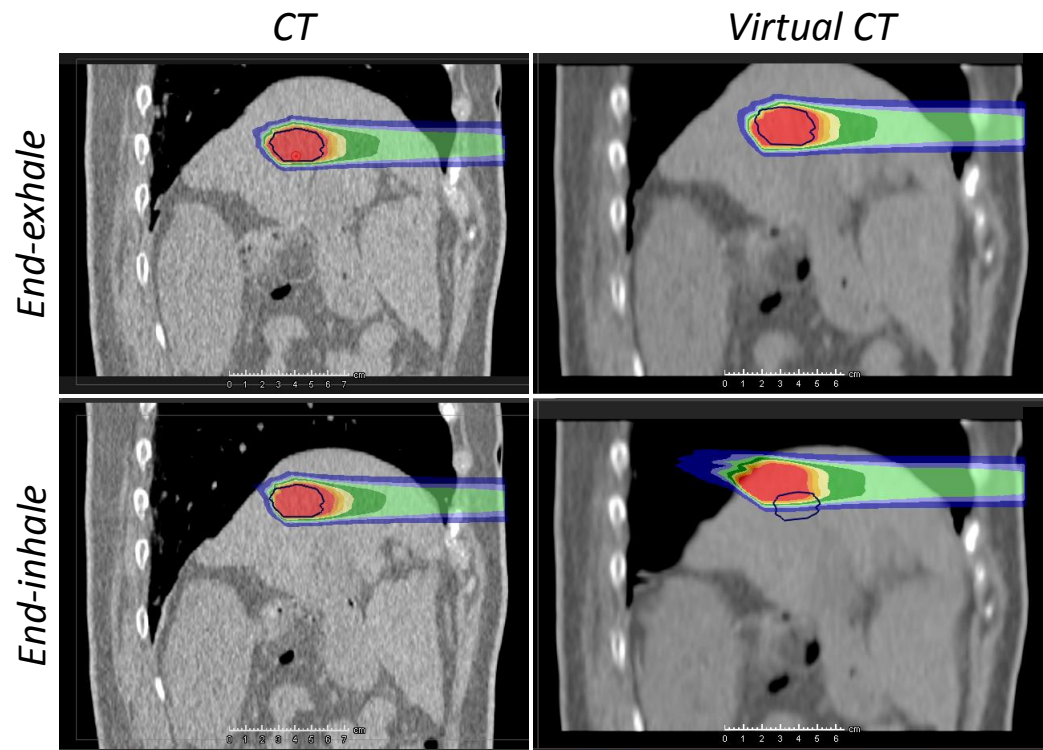
Lung adenocarcinoma, P, 60 Gy(RBE) 10fr/6Gy(RBE), IMPT



# On development (2): use of MRI

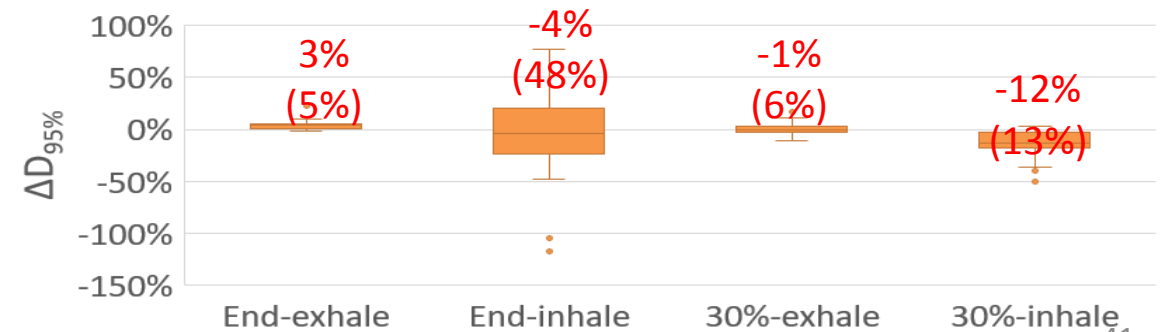


**POLITECNICO**  
MILANO 1863



CIRT plans optimized on end-exhale CT, recomputed on other phases and on virtual 4DCT

*[submitted manuscript]*





# On development (3): delivered dose fraction estimation

The 4DD is the averaged sum of the doses calculated on all N (typically 10) individual phases of a 4D CT scan using the planned fluence without considering the time dependence of the delivery fluence. To calculate the 4DDD, details on the time dependence of the delivery fluence are considered together with changes in anatomy owing to respiratory motion (19, 20, 41). A

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Critical Review

**Consensus Guidelines for Implementing  
Pencil-Beam Scanning Proton Therapy for  
Thoracic Malignancies on Behalf of the PTCOG  
Thoracic and Lymphoma Subcommittee**



- From machine log-file: time point each spot is delivered at which geometric position;
- From gating system: which was the patient anatomy at the delivery of a particular spot



**Retrospective evaluation of dose  
degradation per fraction**

# Four-Dimensional Patient Dose Reconstruction for Scanned Ion Beam Therapy of Moving Liver Tumors

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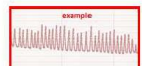
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Volume 89 • Number 1 • 2014

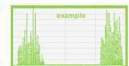
## Log file-based dose reconstruction and accumulation for 4D adaptive pencil beam scanned proton therapy in a clinical treatment planning system: Implementation and proof-of-concept



PBS-PT plan

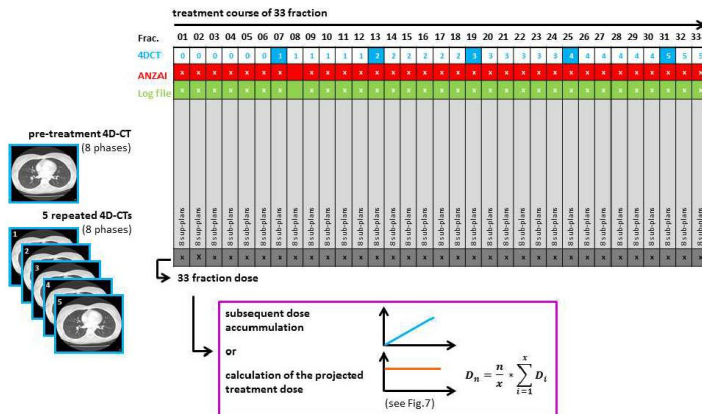


33 different ANZAI traces  
(see Fig.1)



33 delivery log file  
(from a dry-run)

Meijers *et al.*: Med. Phys. 46 (3), March 2019



tionality. For most realistic dose assessment, input data comprised the treatment delivery machine log files of 33 fractions and the patient's breathing patterns, which were acquired during treatment, as well as weekly acquired 4D-CT datasets taken throughout the treatment course. The precision of the 4D dose reconstruction methodology is experimentally validated using a dynamic CIRS thorax phantom.

# Thanks!



.... and questions