

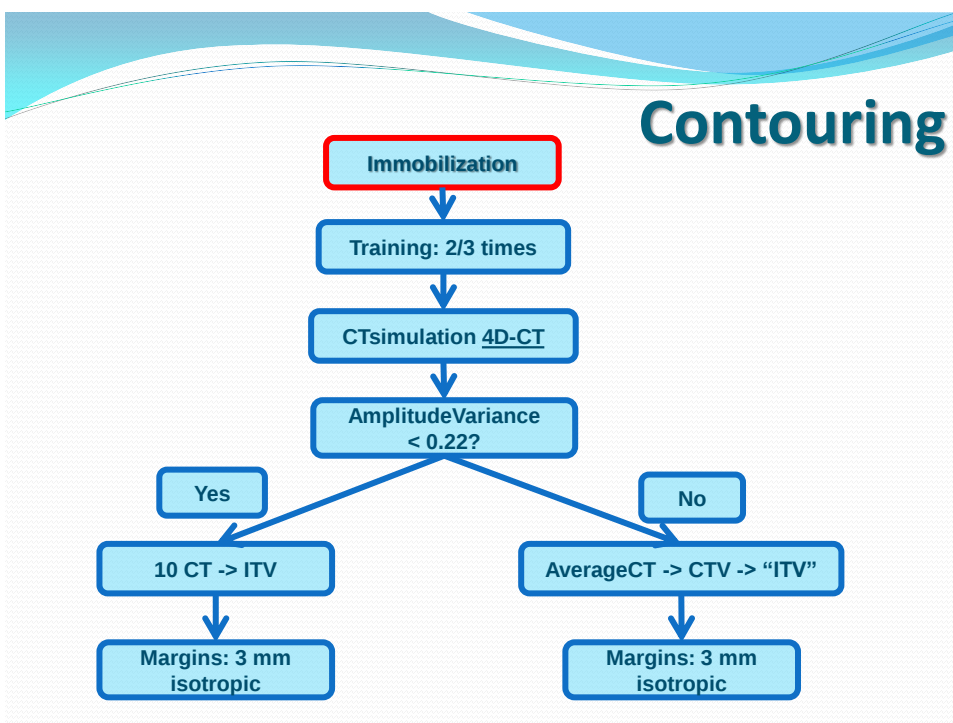
# Lung SBRT : a flowchart for decision making from 4DCT to 4DCBCT

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**24/25 OTTOBRE 2014**  
Università degli studi di Milano  
**STEREOTACTIC  
BODY  
RADIATION  
THERAPY**  
Implementazione, Sostenibilità, Avanzamento Tecnologico  
e Risultati a Confronto

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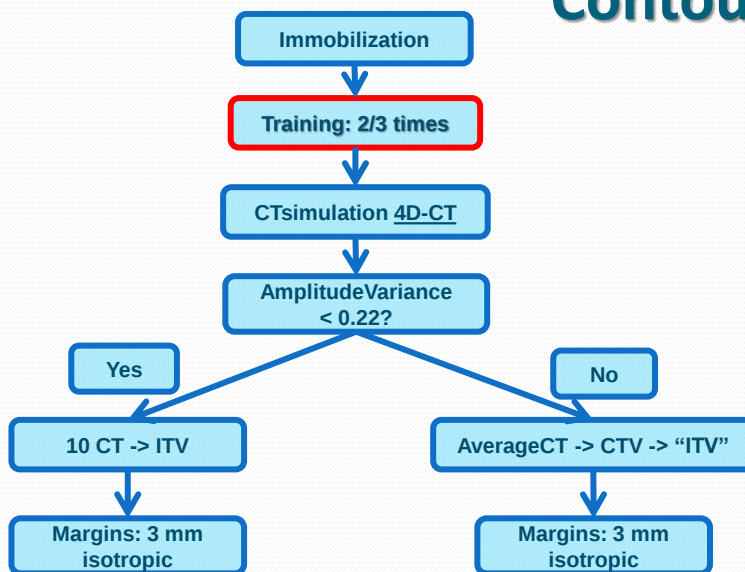


## Immobilization



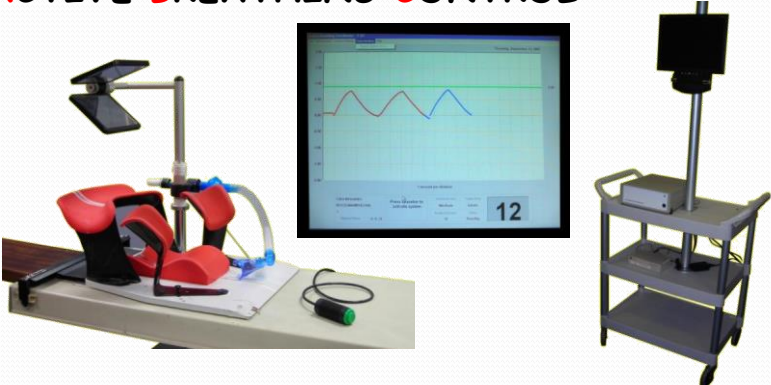
Vacuum immobilisation reduces tumour excursion and minimizes intrafraction error in a cohort study of stereotactic ablative body radiotherapy for pulmonary metastases (Shankar Siva et al 2014)

## Contouring

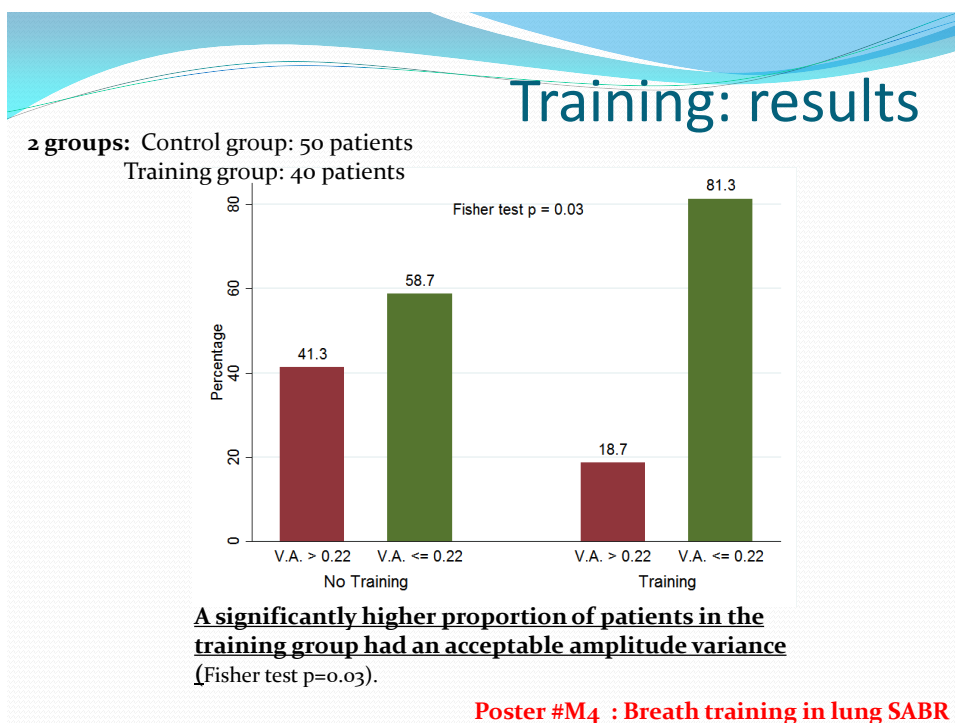


# Training

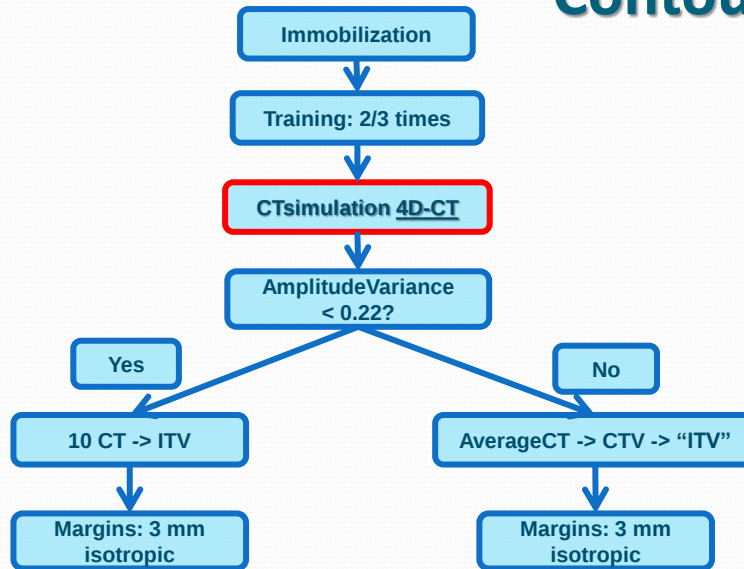
## ACTIVE BREATHING CONTROL



**To reduce variability of  
respiratory movements  
during the radiotherapy treatment**



## Contouring



## CT-Simulation 4D-CT



Lung cancer SBRT

4D-CT-based target volume definition in stereotactic radiotherapy of lung tumours: Comparison with a conventional technique using individual margins

Holger Hof<sup>a,\*</sup>, Bernhard Rhein<sup>b</sup>, Peter Haering<sup>b</sup>, Annette Kopp-Schneider<sup>c</sup>, Jürgen Debus<sup>a</sup>, Klaus Herfarth<sup>a</sup>

<sup>a</sup> University of Heidelberg, Department of Radiation Oncology, Heidelberg, Germany; <sup>b</sup> German Cancer Research Center (DKFZ), Department of Medical Physics, Heidelberg, Germany; <sup>c</sup> German Cancer Research Center (DKFZ), Division of Biostatistics, Heidelberg, Germany

**Results:** Volumetric examinations revealed a significant reduction of the mean PTV by 4D-CT from 57.7 to 40.7 cm<sup>3</sup> (31%) ( $p < 0.001$ ). A significant inverse correlation was found for the motion vector and the amount of inclusion of PTV<sub>4D</sub> in PTV<sub>conv</sub> ( $r = -0.69$ , 90% confidence limits:  $-0.87$  and  $-0.34$ ,  $p = 0.007$ ). Mean lung dose (MLD) was decreased significantly by 17% ( $p < 0.001$ ).

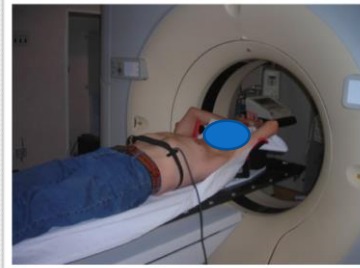
**Conclusions:** In SBRT of lung tumours the mere use of individual margins for target volume definition cannot compensate for the additional effects that the implementation of 4D-CT phases can offer.

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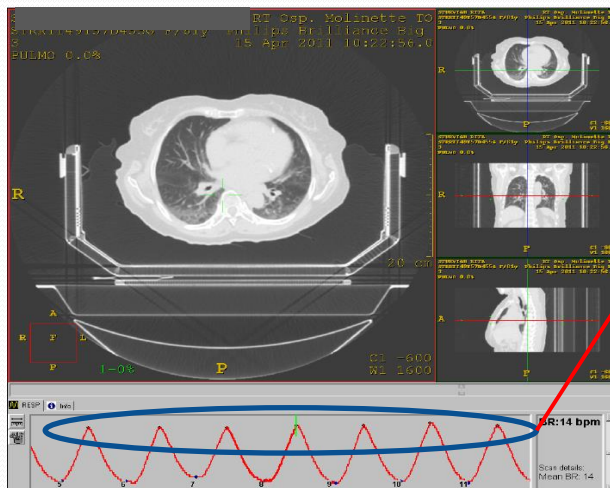
## CT-Simulation 4D-CT



Measurement of abdominal  
diameter



## CT-Simulation 4D-CT



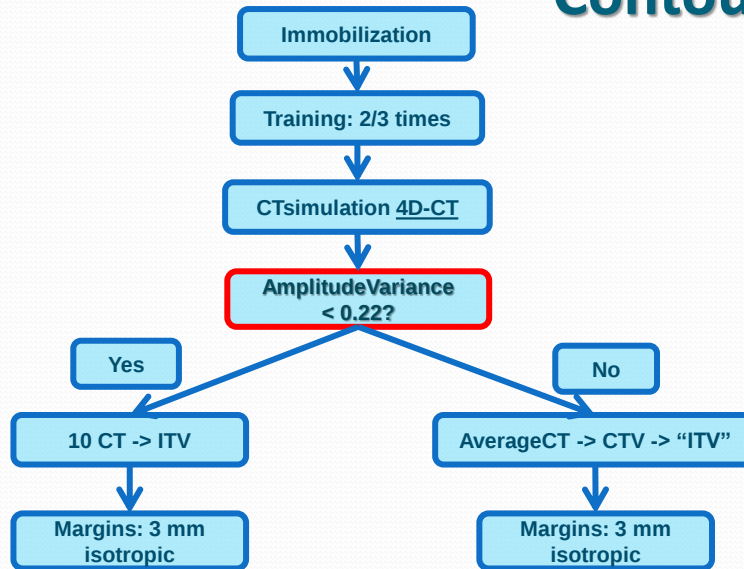
After an analysis of  
amplitude variance:

We choose as suitable  
for an accurate 4D-CT  
the threshold value of  
**20% ( $\pm 10\%$ )**

- **Phase binning:**

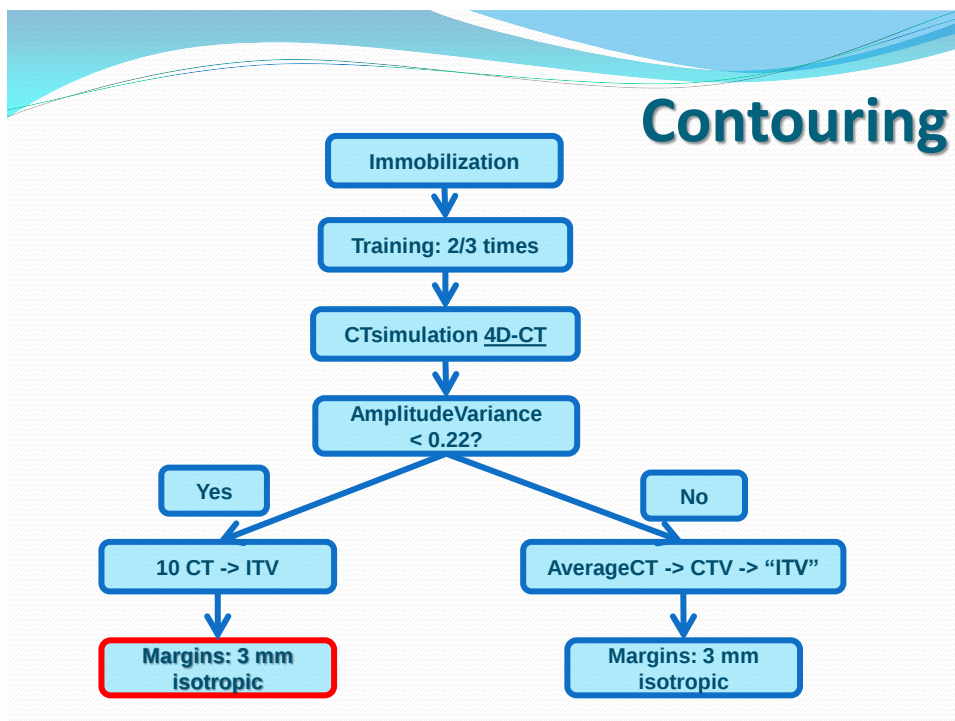
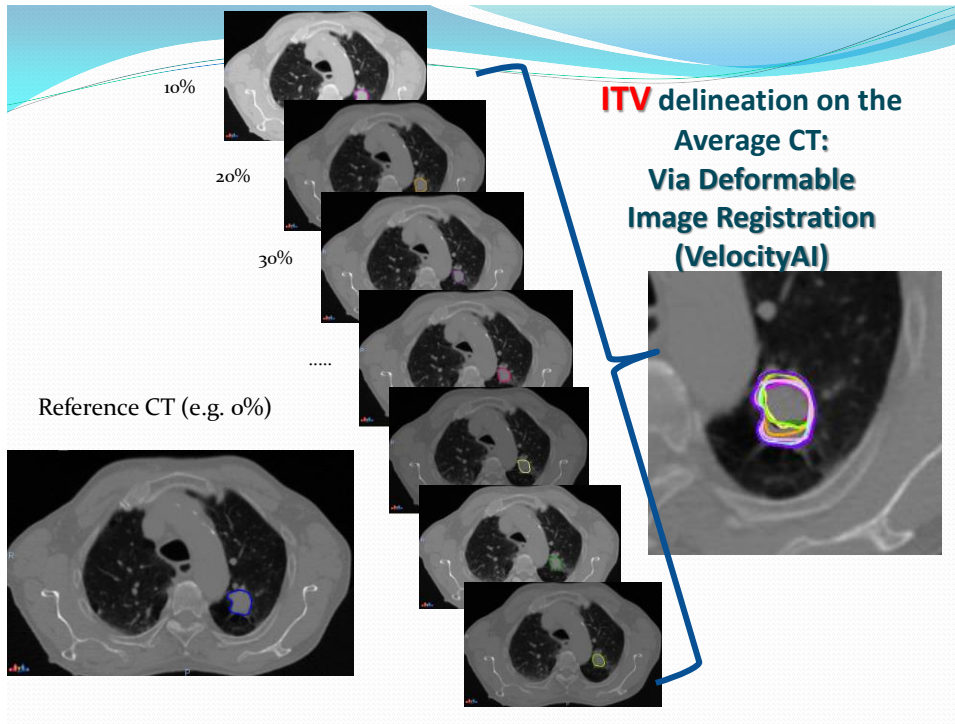
10 series of images corresponding to 10 phases of respiratory cycle

## Contouring



**AV < 0.22**





## Margins

Localization accuracy was quantified by the residual tumor misalignment measured in the second 4D-CBCT scan acquired for validation and expressed in terms of systematic ( $\Sigma$ ), and random ( $\sigma$ ) errors.

For dose prescription at 80% instead of 95%, for a lung target ( $\sigma_p = 0.64$ ):

$$M = 2.5\Sigma + 0.84\sqrt{(\sigma_p^2 + \sigma^2)} - 0.84\sigma_p^2$$

## Margins

We calculated:

LL: 1.6 mm

AP: 1.9 mm

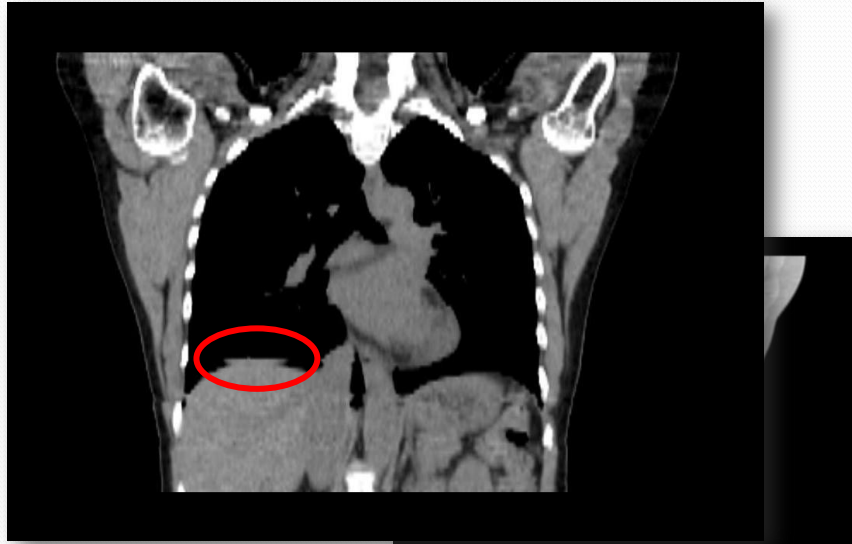
CC: 2.4 mm



We choose:

**3 mm isotropic** (from ITV)

**AV > 0.22**

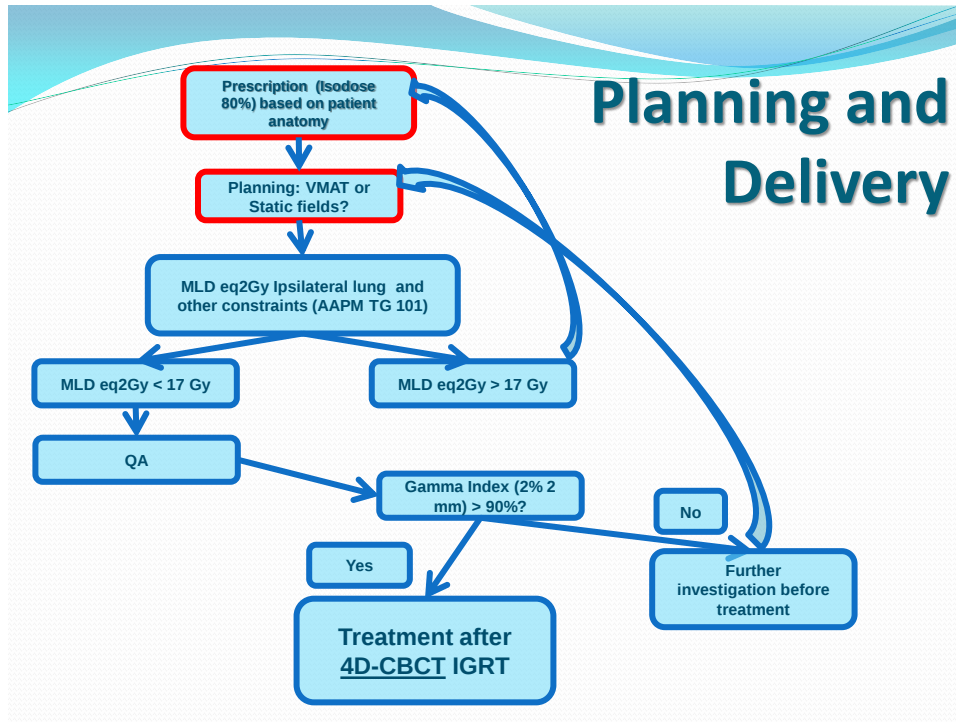


## Contouring: 'statistic ITV'

- Contour of CTV on the Average CT obtained from the 4D-CT scan.
- Obtain a 'statistic ITV' (sITV) by adding margins related to the location of the tumor. We based the amplitude of these margins on a statistical analysis of the tumor motion. Then we used the mean value and 2 SD to define margins for each lobe and for each axis:

|                      |                 |                 |                 |
|----------------------|-----------------|-----------------|-----------------|
| ➤ Inferior lobe(cm): | LL: <b>0.11</b> | AP: <b>0.43</b> | CC: <b>0.65</b> |
| ➤ Medium lobe(cm):   | LL: <b>0.10</b> | AP: <b>0.35</b> | CC: <b>0.60</b> |
| ➤ Superior lobe(cm): | LL: <b>0.10</b> | AP: <b>0.34</b> | CC: <b>0.58</b> |

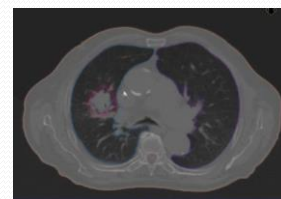
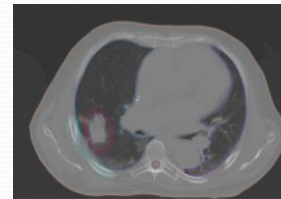
- Then we add **3 mm isotropic** (from sITV) to obtain PTV



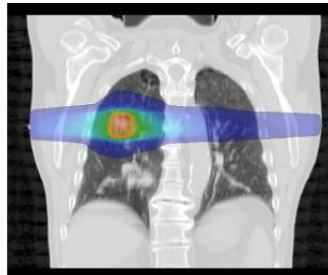
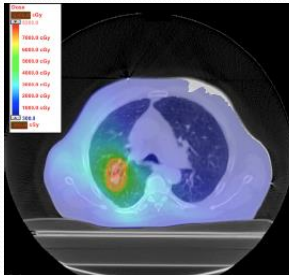
## Prescriptions

### Clinical routine: “risk-adapted” SBRT protocol

- **Peripheral lesions (T1a-T1b):**
  - 45-54 Gy/ 3 fractions
- **Peripheral lesions, with extensive contact with the chest wall, or larger tumors (T2a):**
  - 55 Gy/ 5 fractions
- **Central lesions:**
  - 60 Gy/ 8 fractions

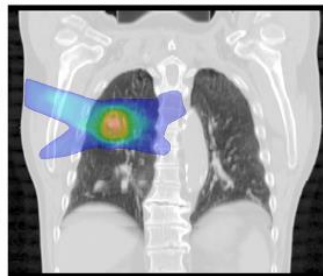
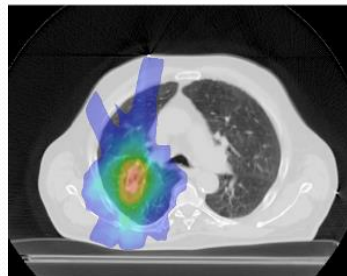


## Step & Shoot or VMAT?

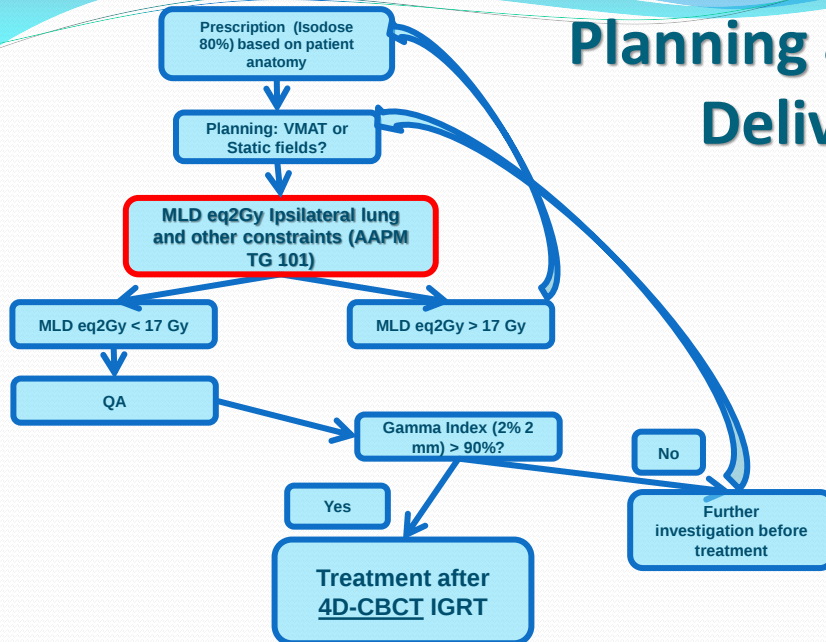


Planning with:  
Elekta CMS  
**Monaco v.3.2/3.3**

Grid calculation 2 mm,  
Montecarlo Variance 1.5%



## Planning and Delivery



## MLD<sub>2Gy</sub> Ipsilateral Lung

Table IV. Logistic regression analysis (correlation with RTOG grade 2-3 pulmonary toxicity)

|                          | Odds Ratio | Std. Err. | z     | p     | 95% Confidence Interval |       |
|--------------------------|------------|-----------|-------|-------|-------------------------|-------|
| MLD <sub>2</sub>         | 1.52       | 0.24      | 2.66  | 0.008 | 1.12                    | 2.08  |
| Primary/ Metastatic      | 3.03       | 3.98      | 0.84  | 0.399 | 0.23                    | 39.79 |
| Central/Peripheral       | 0.54       | 0.70      | -0.48 | 0.634 | 0.04                    | 6.75  |
| Superior/Median/Inferior | 1.53       | 0.89      | 0.73  | 0.464 | 0.49                    | 4.79  |
| Lobe                     |            |           |       |       |                         |       |
| PTV                      | 1.03       | 0.03      | 1.05  | 0.293 | 0.98                    | 1.08  |

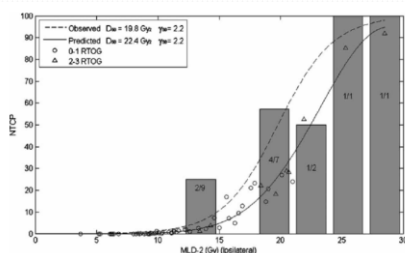


Table III. MLD<sub>2</sub> and NTCP mean values according to RTOG lung toxicity score

|                  | Grade 0-1                     | Grade 2-3                     |
|------------------|-------------------------------|-------------------------------|
| MLD <sub>2</sub> | 11.2 Gy (95% CI 10.1-12.3 Gy) | 20.3 Gy (95% CI 16.6-23.9 Gy) |
| NTCP             | 4% (95% CI 2-5.9%)            | 37% (95% CI 11.6-62.3%)       |

### Dosimetric predictors of radiation-induced lung injury in stereotactic body radiation therapy

*Acta Oncologica*, 2009; 48: 571-577

## MLD<sub>2Gy</sub> Ipsilateral Lung

### Decomposition analysis of differential dose volume histograms

Frank Van den Heuvel Ph.D.<sup>(a)</sup>  
Department of Oncology and Experimental Radiation Therapy  
Universiteit Leuven, Belgium

Dose volume histograms are a common tool to assess the value of a treatment plan for various forms of radiation therapy treatment. The purpose of this work is to introduce, validate, and apply a set of tools to analyze differential dose volume histograms by decomposing them into physically and clinically meaningful normal distributions. A weighted sum of the decomposed normal distributions (e.g. weighted dose) is proposed as a new measure of target dose, rather than the more unstable point dose.

The method and its theory are presented and validated using simulated distributions. Additional validation is performed by analyzing simple four field box techniques encompassing a pre-defined target, using different treatment energies inside a waterphantom.

Furthermore, two clinical situations are analyzed using this methodology to illustrate practical usefulness. A comparison of a treatment plan for a breast patient using a tangential field setup with wedges is compared to a comparable geometry using dose compensators. Finally, a normal tissue complication probability (NTCP) calculation is refined using this decomposition. The NTCP calculation is performed on a liver as organ at risk in a treatment of a mesothelioma patient with involvement of the right lung.

The comparison of the wedged breast treatment versus the compensator technique yields comparable classical dose parameters (e.g. Conformity Index  $\approx 1$  and equal dose at the ICRU dose point). The methodology proposed here shows a 4% difference in weighted dose outlining the difference in treatment using a single parameter instead of at least two in a classical analysis (e.g. mean dose, and maximal dose, or total dose variance). NTCP-calculations for the mesothelioma case are generated automatically and show a 3% decrease with respect to the classical calculation. The decrease is slightly dependant on the fractionation and on the  $\alpha/\beta$ -value utilized.

In conclusion, this method is able to distinguish clinically important differences between treatment plans using a single parameter. This methodology shows promise as an objective tool for analyzing NTCP and doses in larger studies, as the only information needed is the dose volume histogram.

## Constraints

## Stereotactic body radiation therapy: The report of AAPM Task Group 101

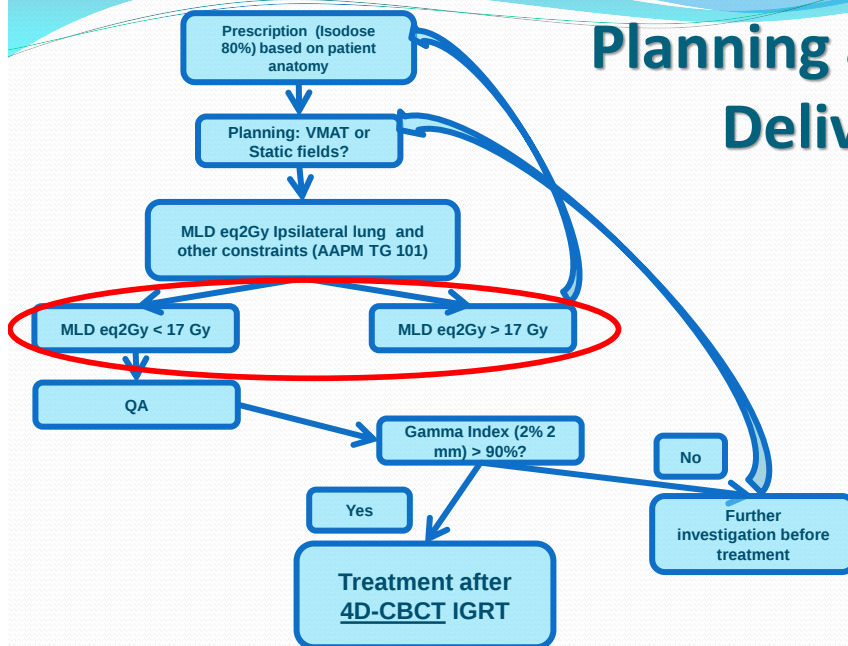
Stanley H. Benedict, Chairman<sup>(a)</sup>

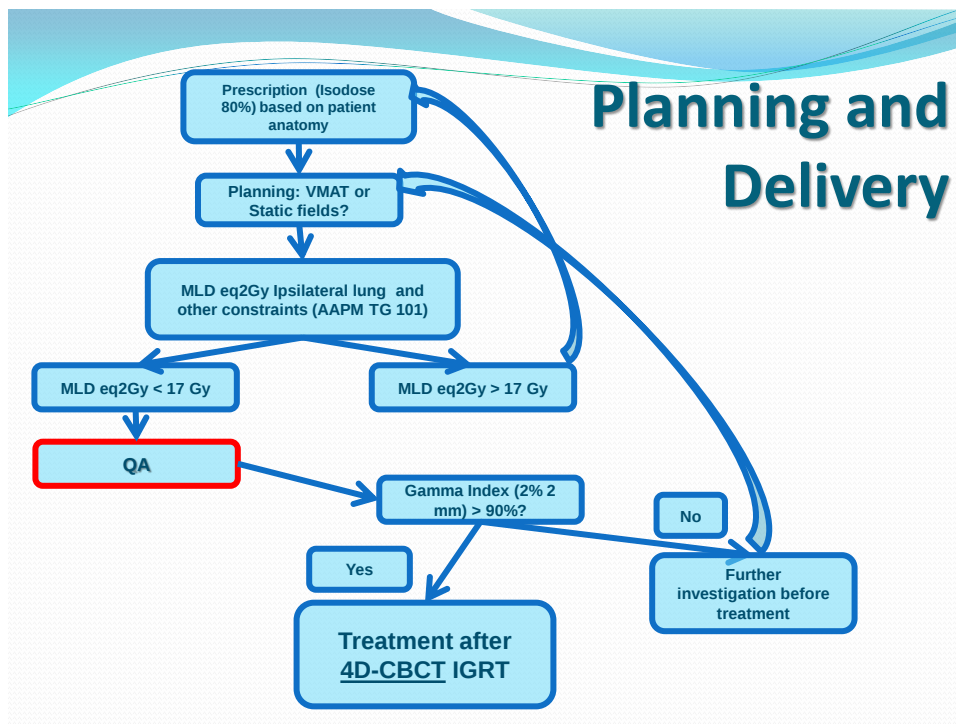
University of Virginia Health System, Charlottesville, Virginia 22908

TABLE III. Summary of suggested dose constraints for various critical organs. Note that for serial tissues, the volume-dose constraints are given in terms of the critical maximum tissue volume that should receive a dose equal to or greater than the indicated threshold dose for the given number of fractions used. For parallel tissues, the volume-dose constraints are based on a critical minimum volume of tissue that should receive a dose equal to or less than the indicated threshold dose for the given number of fractions used.

[illegible]

# Planning and Delivery

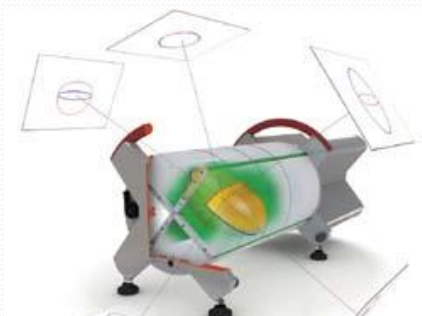
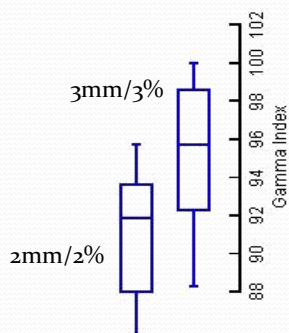


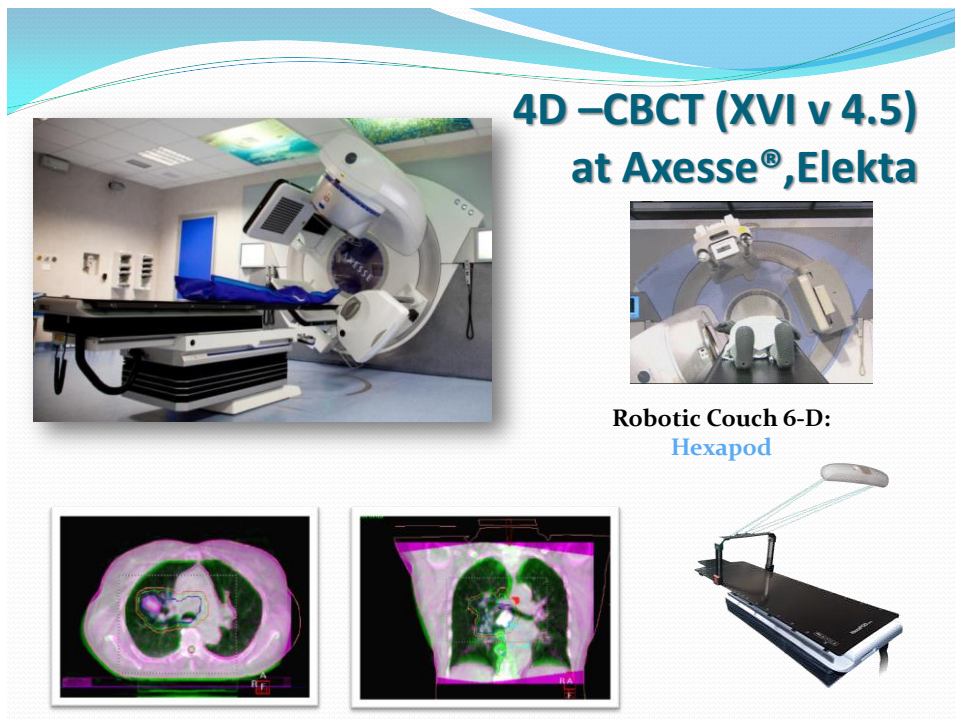
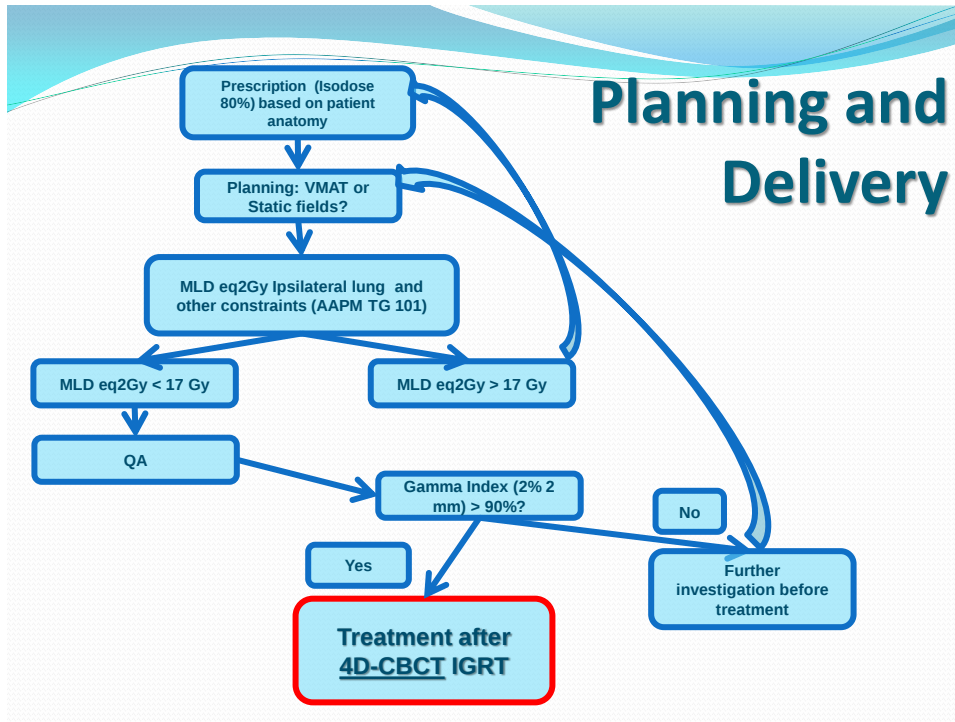


## Patient Specific pre-treatment QA: Delta 4

Acceptance level:

- 1 step: 2mm/2% > 90%
- 2 step 3mm/3% > 95%

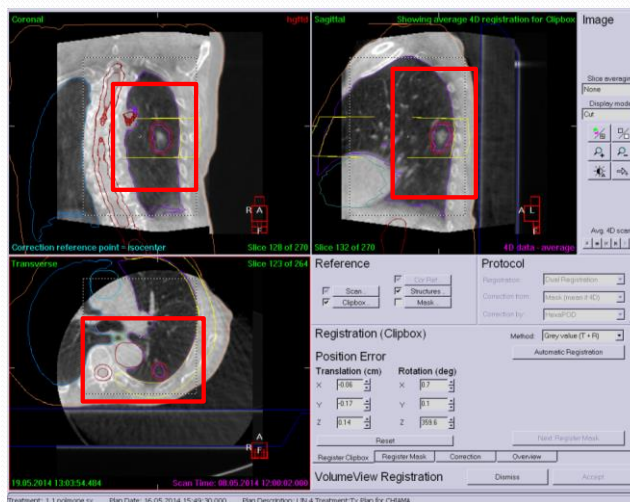




## 4D-CBCT: correction protocol

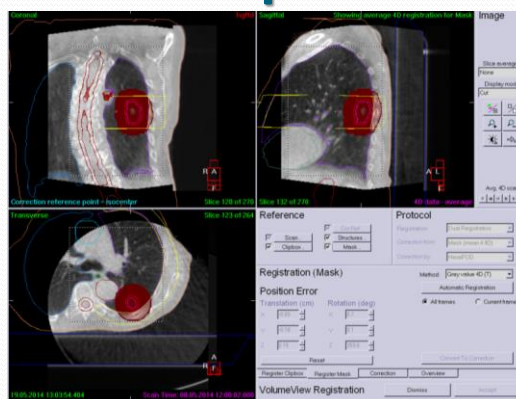
First, the bony anatomy was rigidly registered using a user-defined 3D rectangular-shaped ROI.

We correct set-up of the patient.



## 4D-CBCT: correction protocol

- Second, tumor motion analysis was performed using a local rigid registration, based on a 3D-shaped ROI (PTV expanded by 1.5 cm)
- This ROI was automatically registered (translations only) to each phase of a 4D-CBCT scan yielding the tumor trajectory relative to the planned tumor position.



Rotation and translation corrected by:  
Robotic Couch 6-D: **Hexapod**



## Baseline shift

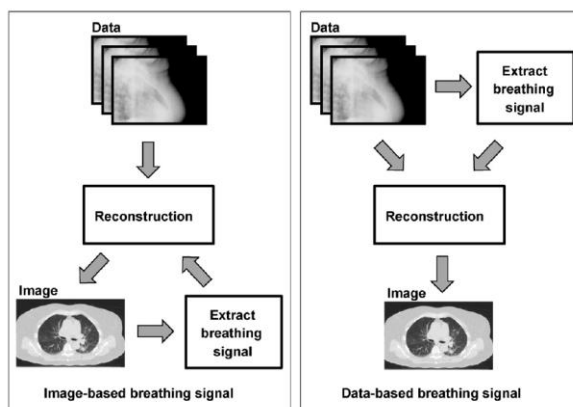
The measured tumor trajectory was averaged (time-weighted) to quantify displacements of the mean tumor position. Corrections for **baseline shifts** were validated.

Abdominal compression was effective for reducing the amplitude of tumor motion. However the use of abdominal compression seemed to increase the **interfraction variation** in tumor position, despite reducing lung tumor motion. The daily tumor position deviated more systematically from the tumor position in the planning CT. Therefore, **target matching is required to correct or minimize the interfraction variation.**



## Grazie dell'attenzione

## Correlate 4D-CT and 4D-CBCT phase binning



Via DIR (VelocityAI) analysis