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# Clinical implementation of **breath-hold** with **nasal high flow therapy** for **photons & protons** at **Maastricht**

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## Device used at Maastrro: AIRVO®

- Delivery of high flow of humidified heated air and supplemental oxygen through a nasal interface
  - **Nasal High Flow therapy (NHFT) (AIRVO®)** to support **voluntary breath hold**
    - Nasal cannula
    - High flow 40L/min (2-60L/min)
    - Heated (31-34-37°C)
    - Insulated tube
    - Humidified
    - Oxygen 21%-80%
  - Easy set-up
  - Combined with surface scanning with visual feedback
- New taxonomy: HFNC



# Use of NHFT in non-RT setting

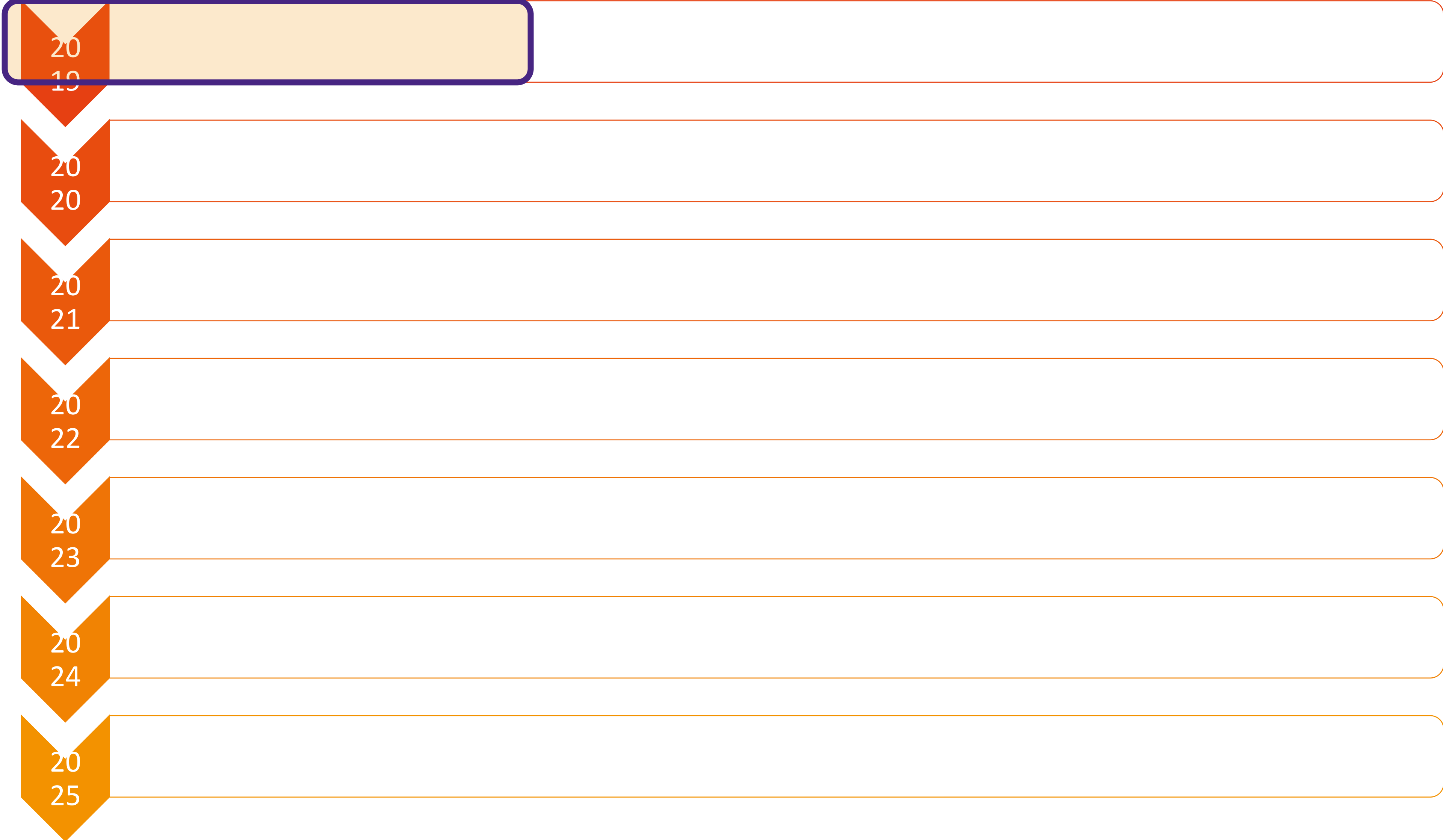
Is used in hospitals, but also at home

Website vendor →

## Recommendations for NHF use

Clinical applications/ scenarios	ESICM <sup>2,3</sup>	ACP <sup>1</sup>	SSC <sup>3</sup>	AARC <sup>4</sup>	ERS <sup>5</sup>	TSANZ <sup>6</sup>	JSICM JRS JSRCM <sup>7</sup>	WHO <sup>8</sup>	SRLF SFMU <sup>9</sup>	ACEP <sup>11</sup>	NICE <sup>12</sup>	GOLD <sup>13</sup>
<b>PRIMARY SUPPORT - MEDICAL</b>												
Undifferentiated or moderate respiratory distress										[a]		
Acute hypoxemic respiratory failure (AHRF)	[b,c]	[d]			[e]	[f]			[g]			
Respiratory failure due to pneumonia											[h]	
Acute respiratory distress syndrome (ARDS)	[c]						[i]					
Acute hypercapnic respiratory failure					[j]							
Exacerbation of COPD												[k]
Sepsis-induced AHRF			[l]									
Patients with severe or critical COVID-19	[c]							[m]				
Patients who are immuno-compromised with AHRF				[n]								
<b>PRIMARY SUPPORT - SURGICAL</b>												
Low risk post-operative patients					[o]							
High risk post-operative patients					[p]							
High risk post-cardiothoracic surgery patients	[q]											
<b>PRE-ESCALATION SUPPORT</b>												
Pre-oxygenation for peri-intubation patients	[r]											
<b>DE-ESCALATION SUPPORT</b>												
Post-extubation	[s]	[t]		[u]								
Low risk, extubated patients					[v]							
High risk, extubated patients	[w]				[x]							
<b>COMPLEMENTARY RESPIRATORY SUPPORT</b>												
Rest breaks off NIV					[y]							
<b>PROPHYLACTIC RESPIRATORY SUPPORT</b>												
Patients requiring supplemental oxygen for any reason				[z]								

# NHFT in radiotherapy under breath hold: Maastrro experience



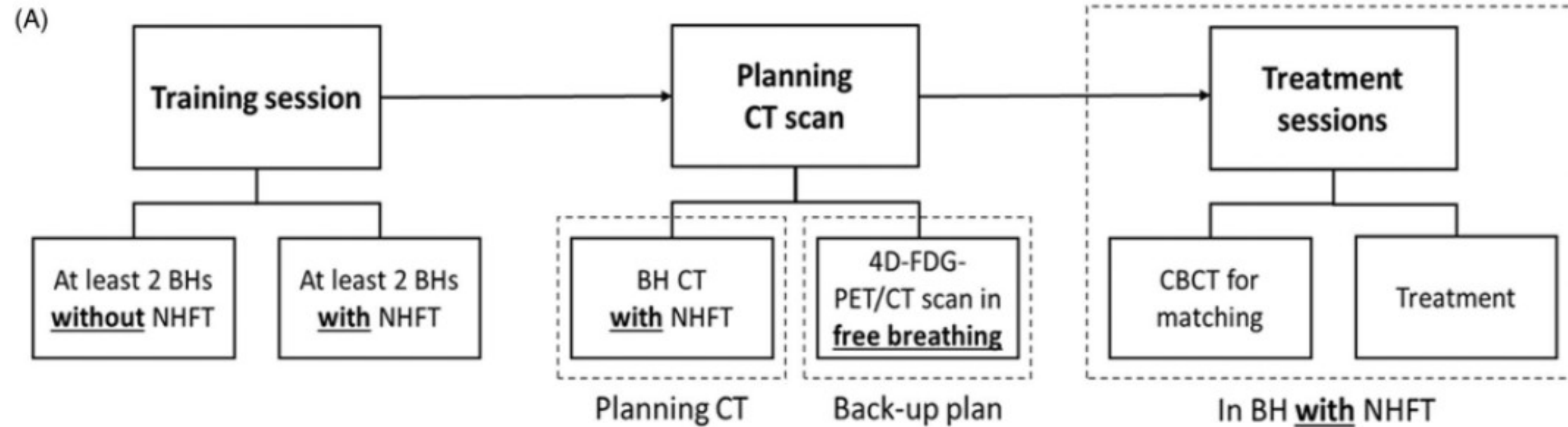
# Lung cancer, photons

## First experience: ENTheR study

Treated with NHFT

- with 80% O<sub>2</sub>
- Flow 40 L/min

Inclusion	10 consecutive locally advanced NSCLC patients undergoing radical <u>photon</u> RT
Methods	Prospective, NHFT, surface scanning, CO <sub>2</sub> -meter, 2019
Endpoints	Tolerability of the treatment <b>Breath hold length</b> Stability & reproducibility (Surface scanning) Subjective tolerance Dose difference between standard treatment plan and treatment plan in BH with NHFT.



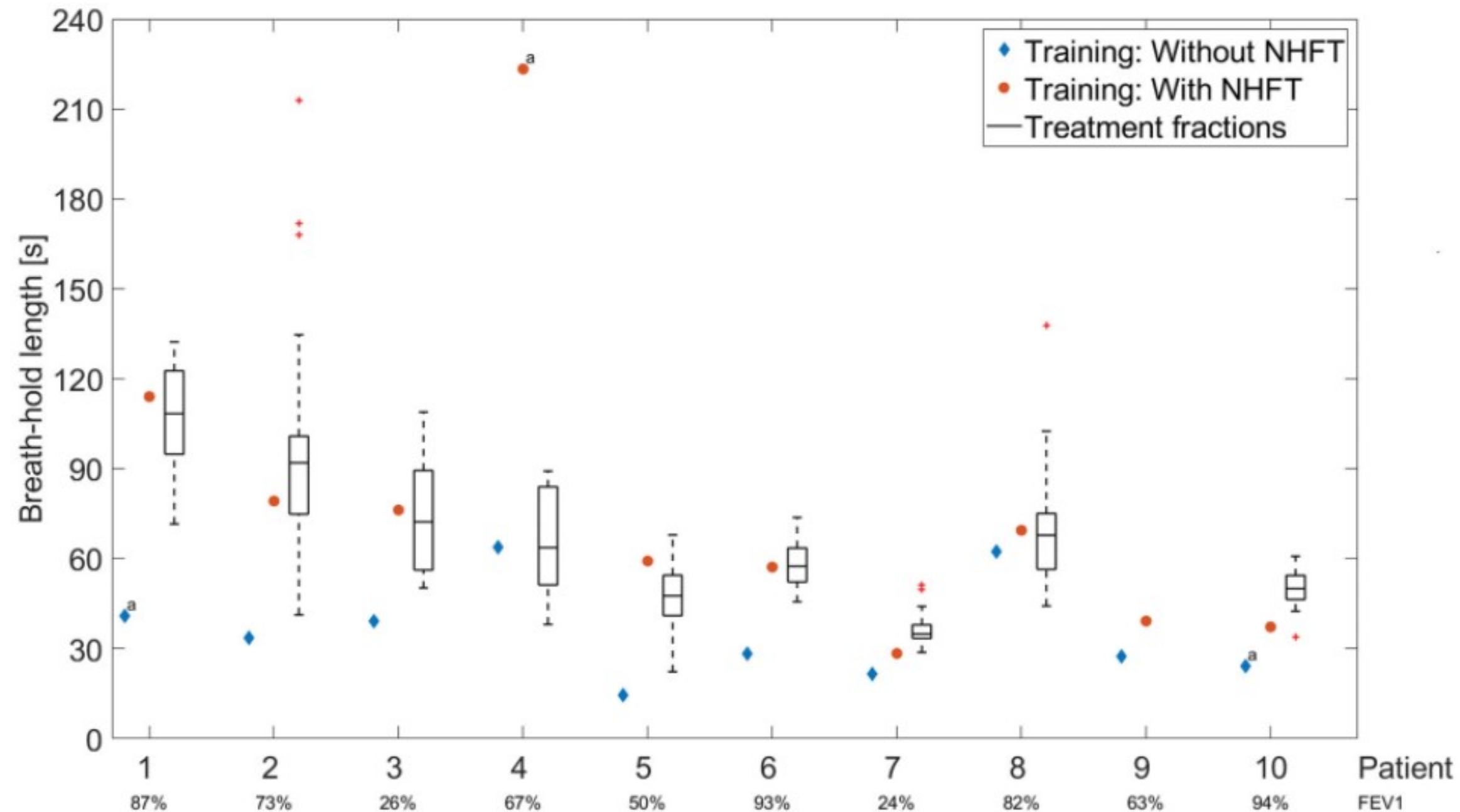
## Visually guided inspiration breath-hold facilitated with nasal high flow therapy in locally advanced lung cancer

Stephanie T. H. Peeters , Femke Vaassen , Colien Hazelaar , Ana Vaniqui , Eva Rousch , Debby Tissen , Esther Van Enkevort , Michiel De Wolf , Michel C. Öllers , Wouter van Elmpt , Karolien Verhoeven , Judith G. M. Van Loon , Bettine A. Vosse , Dirk K. M. De Ruyscher & Gloria Vilches-Freixas

# Lung cancer, photons

## First experience: ENTheR study

Results	9/10 included patients treated within study
	7/9 completed treatment as planned (2 pts had 2 fractions in FB)
	Training session: mean BH duration (p=0.005) <ul style="list-style-type: none"> <li>- Without NHFT: 39 s (range 15-86)</li> <li>- With NHFT: 78 s (range 29-223)</li> <li>- Factor 2.1 (range 1.1-3.9)</li> </ul>
	Intra- and inter-BH variability were both minimal, with submillimeter variability
	Mean inter-fraction tumor position variability: 1.8mm (-1.1–8.1;SD 2.4)

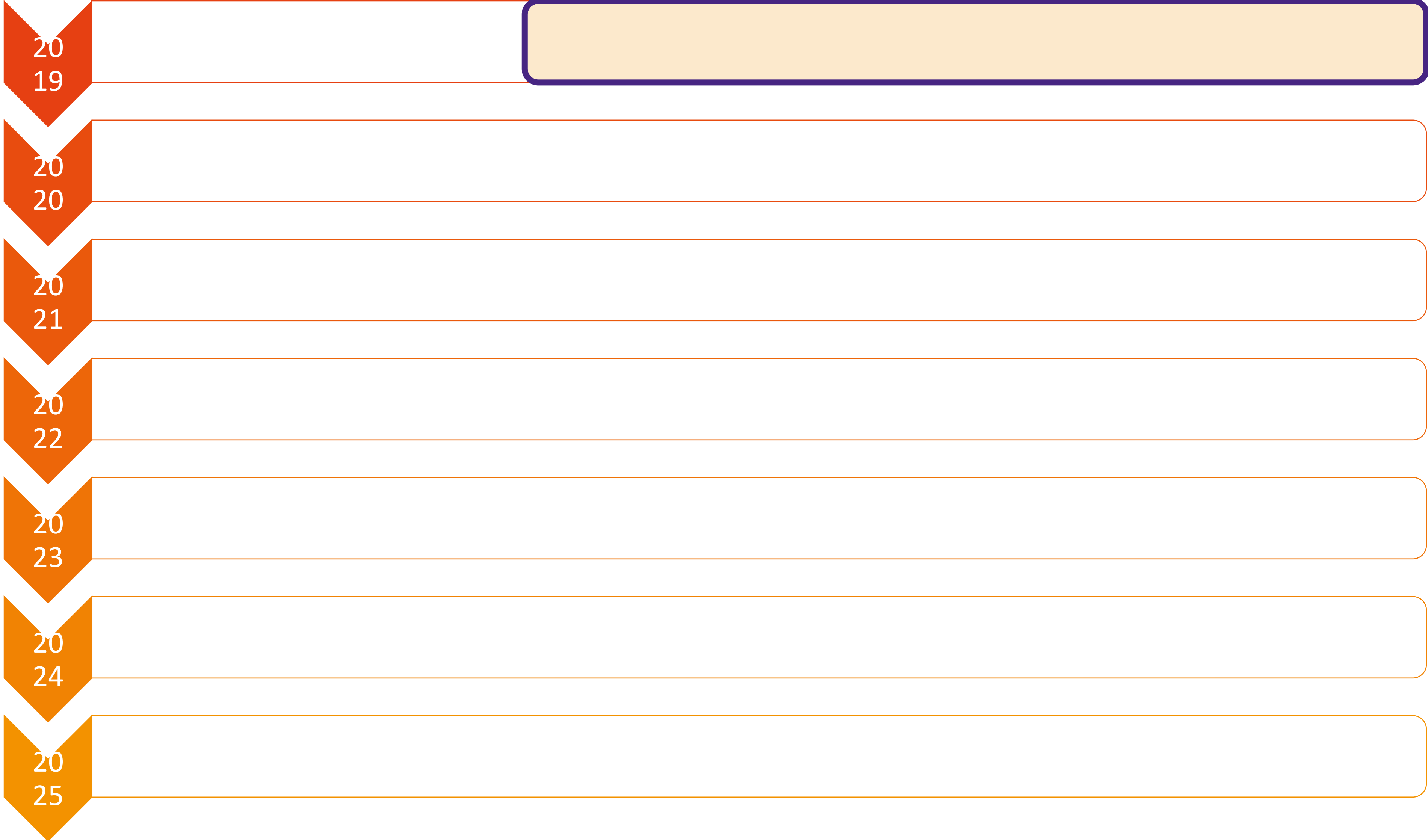


**Table 1.** Patients' characteristics.

	N = 10
Mean age (years) (range)	67 (60–74)
Gender	
Male	6
Female	4
WHO performance status	
0	3
1	5
2	2
Stage (TNM 8)	
Stage IIIA	3
Stage IIIB	3
Stage IIIC	2
Stage IVA	1
Stage IVb	1
Primary tumor location	
Upper lobe	8
Middle lobe	0
Lower lobe	2
FEV1 (%pred): mean (range)	66 (24–94)
Smoking status	
Never smoker	0
Former smoker	6
Active smoker	4
Treatment (n = 9) <sup>a</sup>	
Concurrent chemoradiotherapy	6
Sequential chemoradiotherapy	2
Radiotherapy alone	1
Lung volume (n = 9) <sup>a</sup>	
Free breathing (L) (mean) (range)	4.1 (2.1–5.7)
Breath hold (L) (mean) (range)	6.1 (4.1–8.0)
Relative increase (%) (mean) (range)	150 (114–200)

FEV1: Forced expiratory volume in 1 s.  
<sup>a</sup>1 patient was upstaged to stage IVc at planningCT and was not treated with RT.

# NHFT in radiotherapy under breath hold: Maastrro experience



# Lymphoma, photons

11 mediastinal lymphoma pts included

Treated with photons in BH: 6 with NHFT (pts 1-6), 5 without NHFT (pts 7-11)

Flow of 40 L/min, with 80% O<sub>2</sub>

	NHFT (n)	No NHFT (n)
Number of patients	6	5
Age Median (range) (years)	36 (21–54)	22 (19–33)
Gender		
Male	4	3
Female	2	2
Cardiac risk factors		
Yes	3	1
No	3	4
Stage		
IIA	5	5
IIB	1	0
Histology		
Classic Hodgkin lymphoma	6	5
Chemotherapy regimen		
A(B)VD courses		
2	1 (+2x escBEACOP)	1
3	2	0
4	2	0
5	0	3
Other chemotherapy regimen	1 (3x GDP + BEAM)	1 (reduced dose)
Fractionation schedule		
10x2Gy	2	1
15x2Gy	3	4
20x2Gy	1	0
Extra mediastinal involvement		
Axilla	2	2
Neck	5	4
Periclavicular	5	2
Internal mammary chain	1	0

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Original Article

Radiotherapy for mediastinal lymphoma in breath hold using surface monitoring and nasal high flow oxygen: Clinical experiences and breath hold stability

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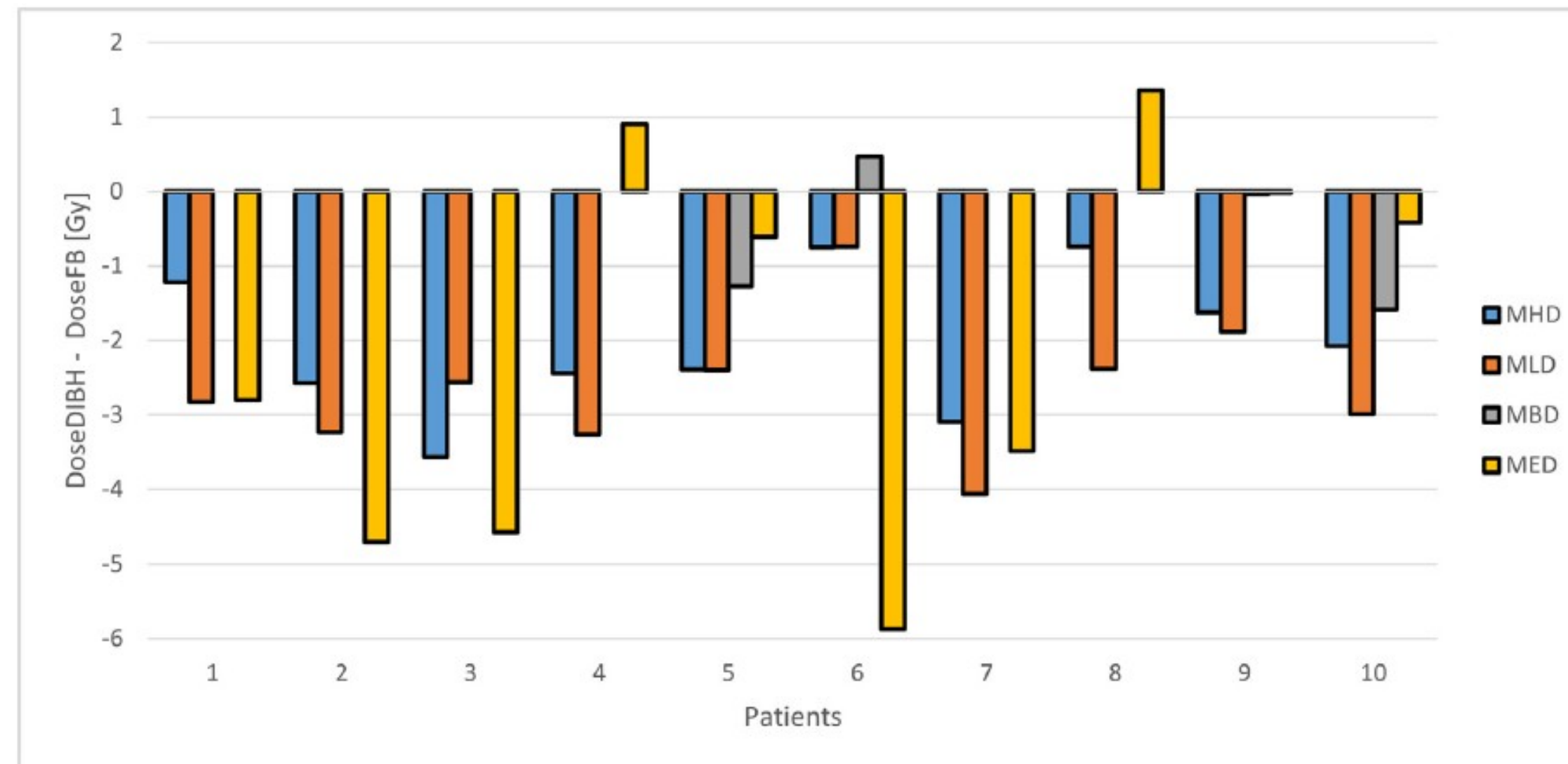


Fig. 4. Dose in breath hold plan minus dose in free breathing plan for mean dose in heart (MHD), lung (MLD), breast (MBD), and esophagus (MED).

# Lymphoma, photons

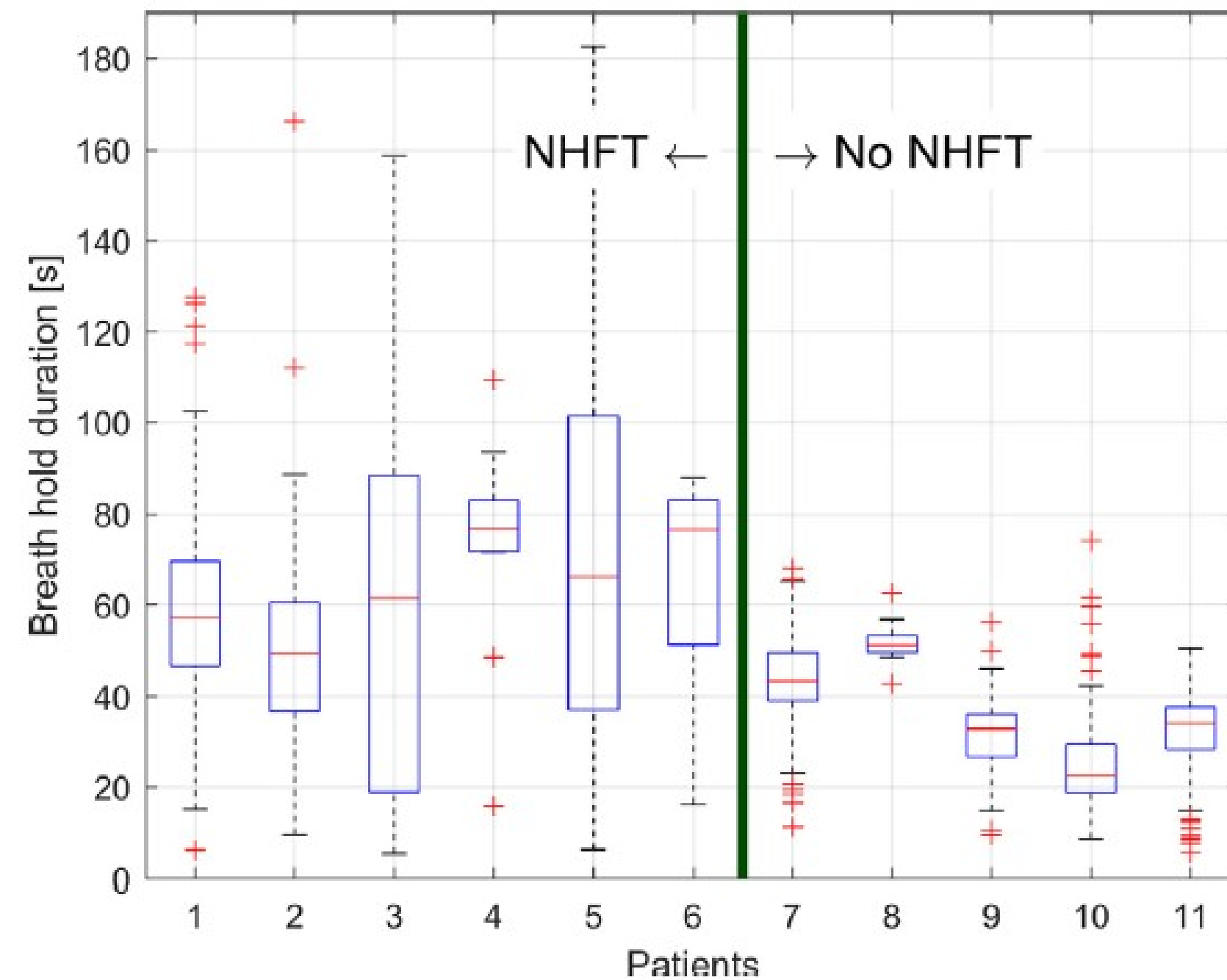
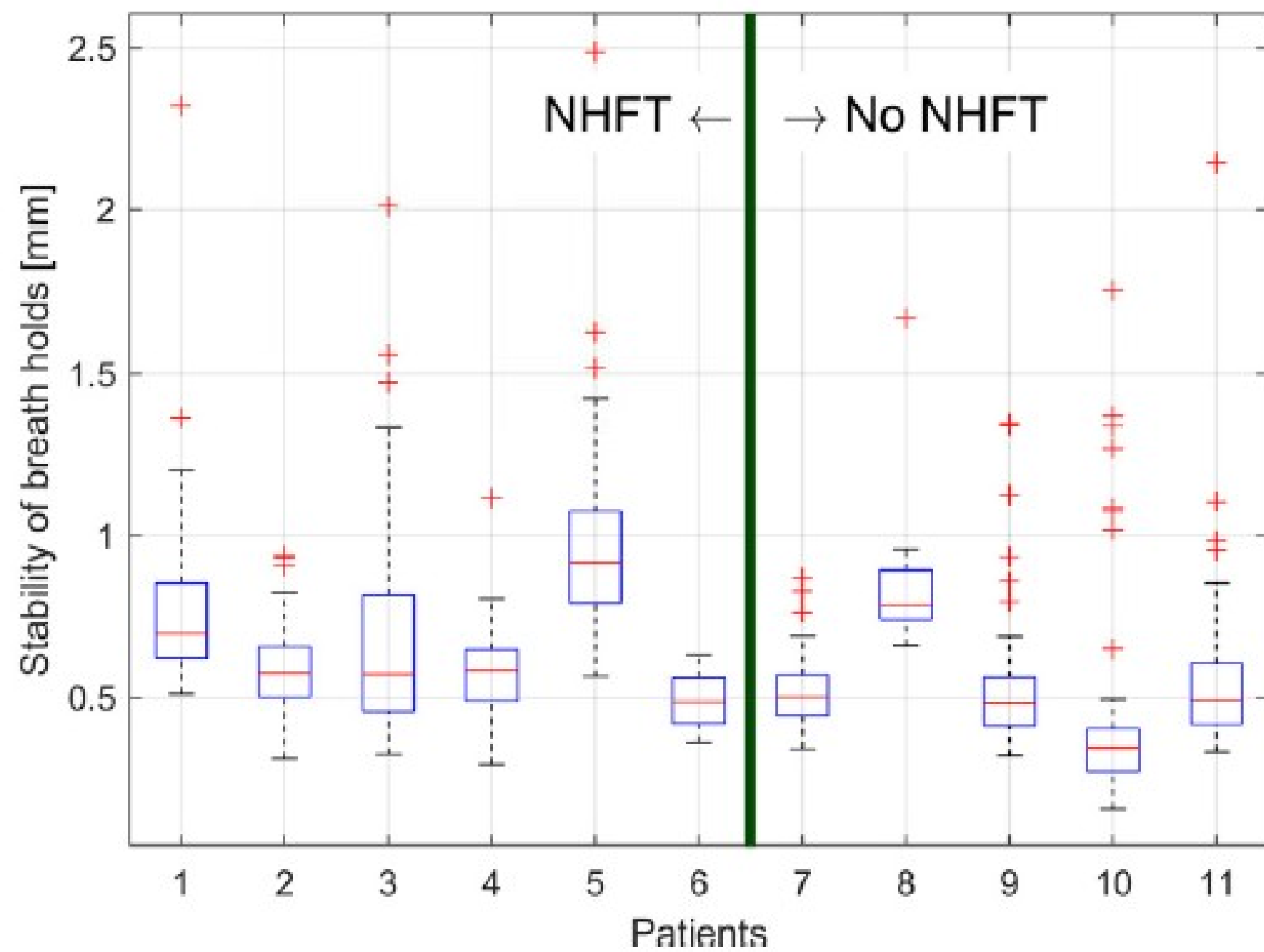
Breath holds were stable

Similar stability with and without NHFT

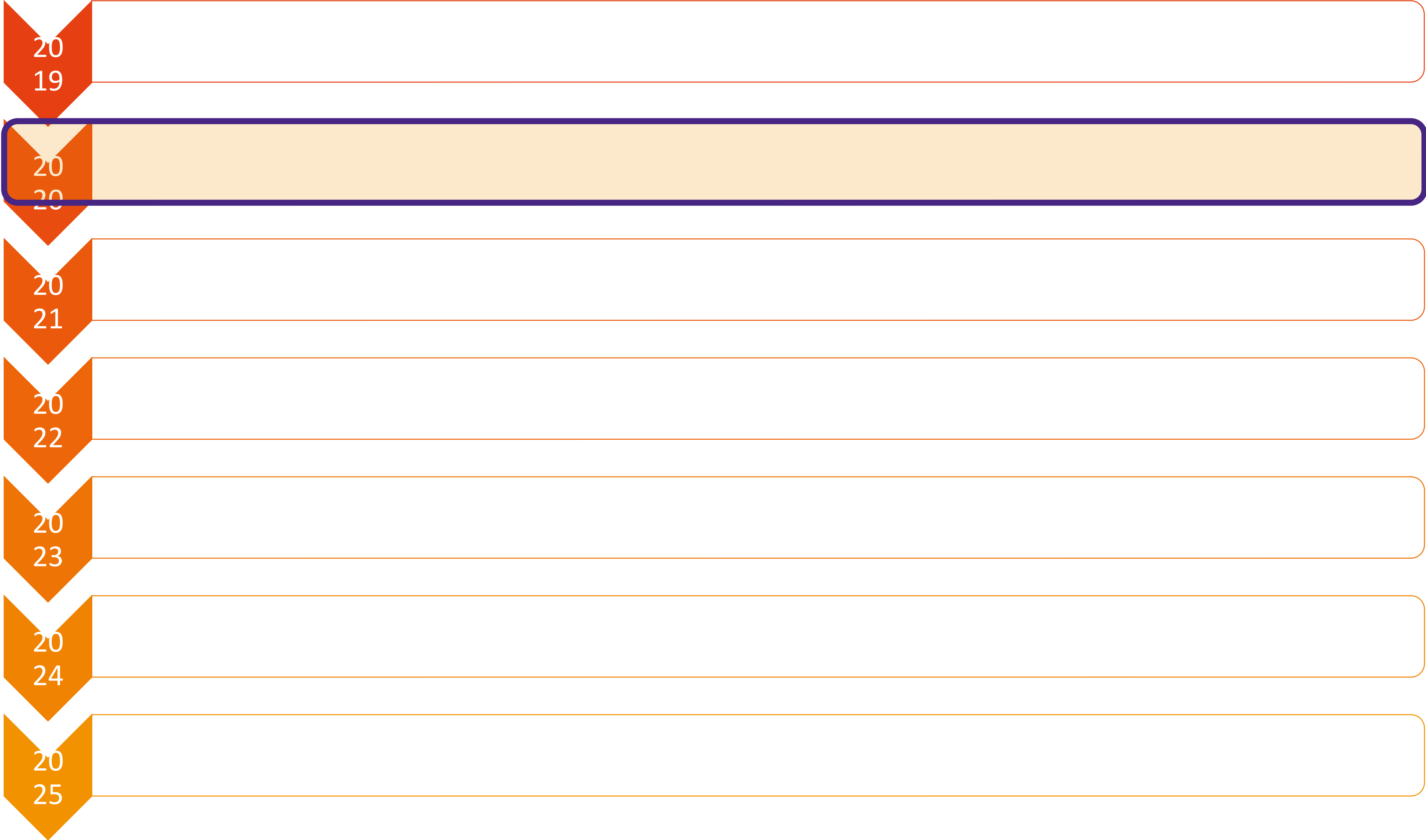
NHFT +/- doubled BH duration

**Table 2**  
Overview of inter and intra fraction random ( $\sigma$ ) and systematic ( $\Sigma$ ) setup errors in left-right (LR), craniocaudal (CC), and anterior-posterior (AP) direction as well as the resulting margins, both for free breathing as well as breath hold.

	Inter fraction motion DIBH [mm]	Intra fraction motion DIBH [mm]	Inter fraction motion free breathing [mm]
$\Sigma_{LR}$	0.9	0.8	0.7
$\Sigma_{CC}$	1.3	0.7	2.1
$\Sigma_{AP}$	1.1	0.5	1
$\sigma_{LR}$	1.2	1.0	0.6
$\sigma_{CC}$	1.9	1.7	1.8
$\sigma_{AP}$	1.3	1.2	1.1
<b>Margins [mm]</b>			
LR	4		3
CC	5		7
AP	4		4



# NHFT in radiotherapy under breath hold: Maastrro experience



# Liver SBRT

- 11 pts treated for liver mets with SBRT

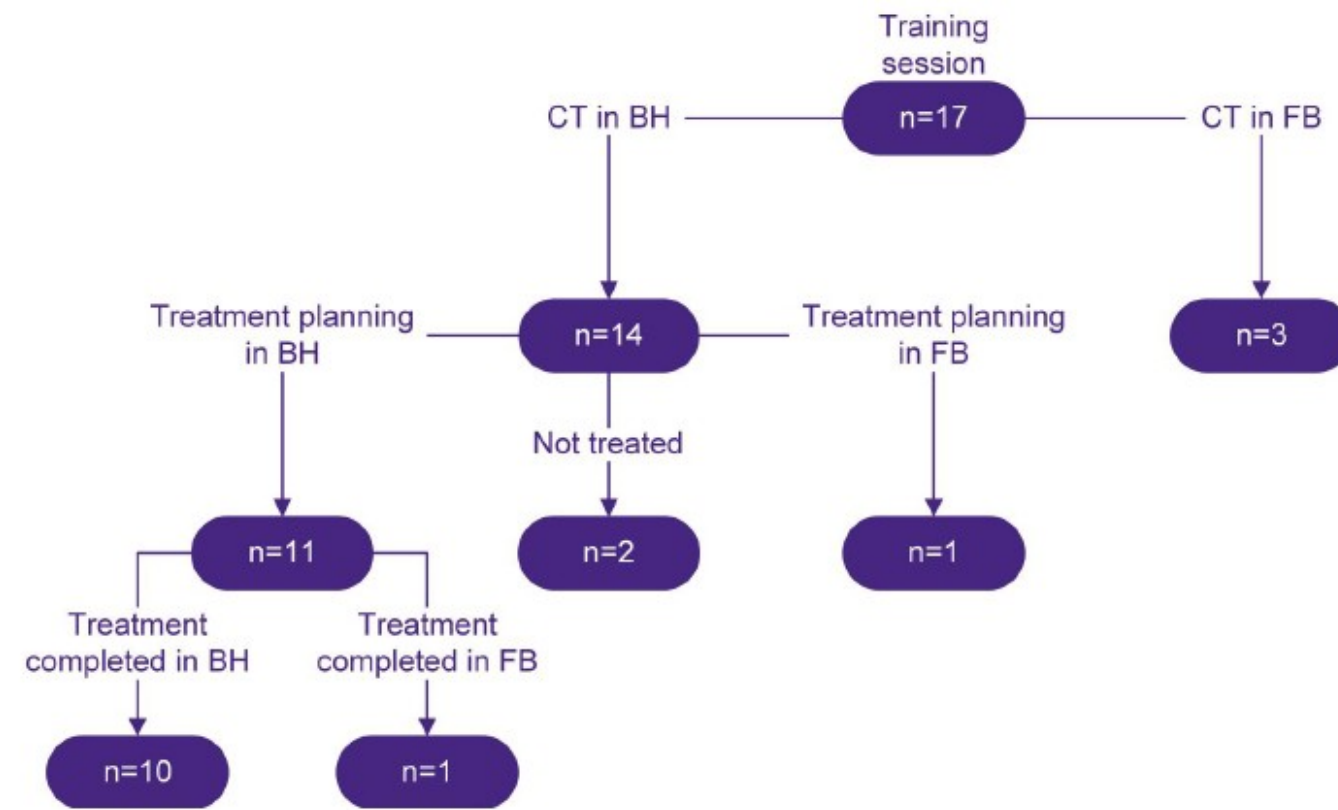


Figure 3. Flowchart showing the number of breath-hold (BH) patients and patients who continued in free-breathing (FB) for each step of the workflow.

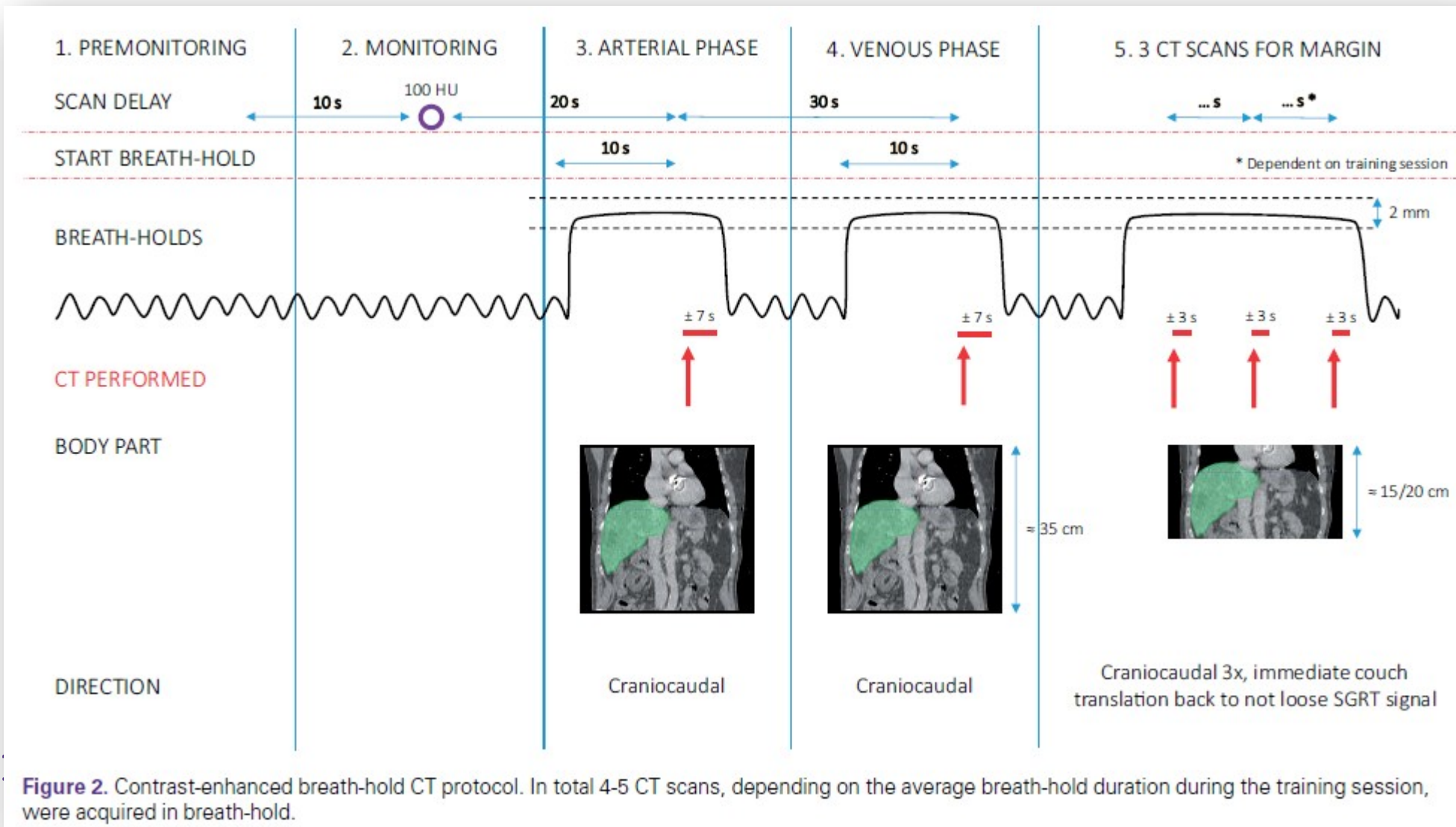

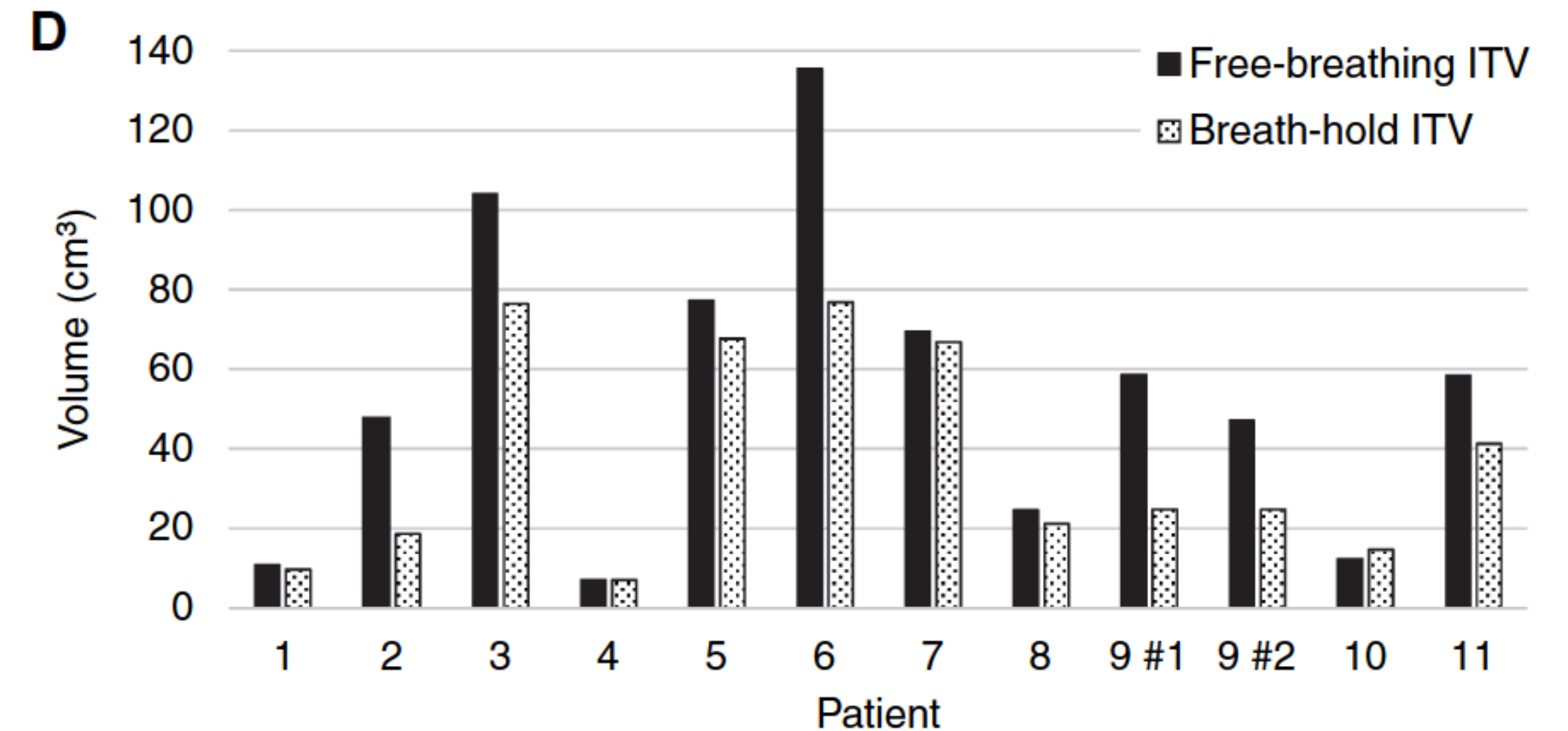


Figure 2. Contrast-enhanced breath-hold CT protocol. In total 4-5 CT scans, depending on the average breath-hold duration during the training session, were acquired in breath-hold.

## Clinical implementation and evaluation of stereotactic liver radiotherapy in inspiration breath-hold using nasal high-flow therapy and surface guidance

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# Liver SBRT

Table 1. Patient and treatment characteristics.

Patient	Sex	Age (y)	PTV (cm <sup>3</sup> )	Dose prescription (fractions × Gy)	Energy	Maximum dose rate (MU/min)	MU/arc	Median number of BHs performed during CBCT imaging (range)	Median number of BHs performed during treatment (range)
1	F	55	58.7	5 × 12	6X-FFF	1400	965/945/865	1 (1-5) <sup>f</sup>	3 (2-8)
2	F	80	95.5	3 × 20	10X-FFF	1600	1832/1614/1514	1 (1-2)	7 (5-10)
3	M	73	177.4	8 × 7.5	10X-FFF	2400	1000/1165 <sup>d</sup> 960/1118	2 (1-4)	6.5 (3-19)
4 <sup>a</sup>	F	72	38.9	5 × 12	10X-FFF <sup>d</sup> 6X-FFF	2400 <sup>d</sup> 1400	1936/1912 <sup>d</sup> 1312/1717	1 (1-2)	6 (4-7)
5	M	73	182.4	5 × 12	6X-FFF	1400	1325/1524	2.5 (1-4)	19 <sup>g</sup>
6	M	67	190.9	5 × 12	6X-FFF	1400	1218/1497	2 (1-3)	8 (4-13)
7 <sup>a</sup>	M	80	242.8	8 × 7.5	6X-FFF	1400	817/859	2 (1-4)	4 (3-8)
8	M	68	91.2 18.5 <sup>b</sup>	5 × 12	6X-FFF	1400	1776/1824	1 (1-5)	5 (3-5)
9	M	76	225.2 <sup>c</sup>	3 × 8	6X-FFF	1400	1232/1136	2 (1-4)	5 (4-14)
10	M	32	72.9	3 × 20	6X-FFF	1400	3065/3065	1.5 (1-4)	9 (6-17)
11	M	70	151.7	3 × 8	6X-FFF	1400	313/783/723/305 <sup>c</sup>	2.5 (1-3)	5 (4-6)

Abbreviations: PTV=planning target volume, MU=monitor unit, BH=breath-hold, CBCT=cone-beam CT, F=female, M=male, FFF=flattening filter free.  
<sup>a</sup>These patients did not have a metastasis in the liver itself but close to the liver: patient 4 had a metastasis next to/below the liver, and patient 7 between the liver and thoracic wall.  
<sup>b</sup>Two PTVs treated in a single plan, of which one was located within the liver and one between the ribs.  
<sup>c</sup>This PTV consisted of multiple GTVs.  
<sup>d</sup>After the first fraction, these patients were treated using a new treatment plan.  
<sup>e</sup>This patient was positioned with the arms alongside the body. More treatment arcs were used to avoid irradiation through the arms.  
<sup>f</sup>CBCT information of one fraction was missing and is therefore not included.  
<sup>g</sup>This patient received only one fraction in breath-hold.

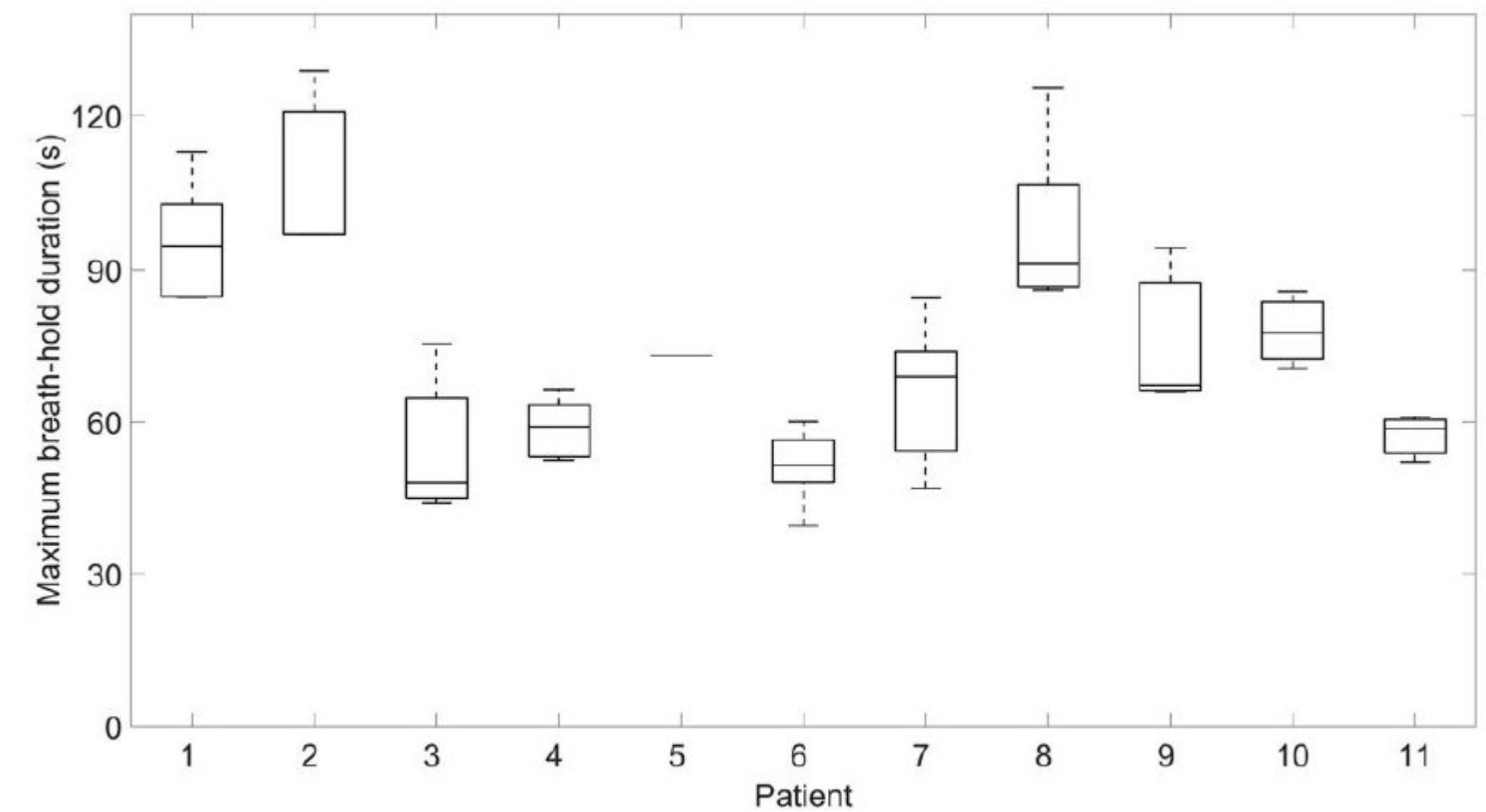
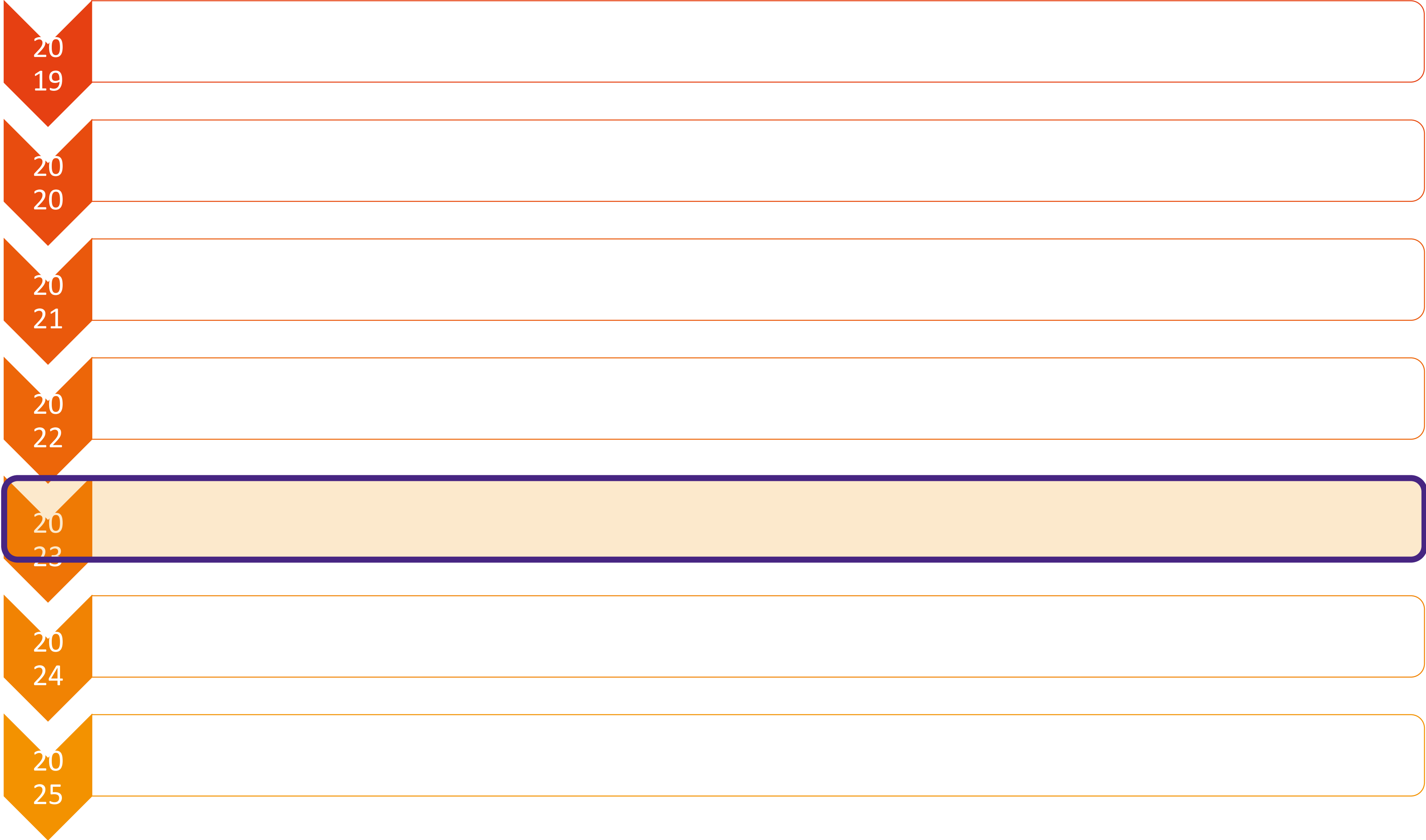


Figure 4. Boxplot showing for each patient the longest breath-hold per fraction during the treatment phase.

# NHFT in radiotherapy under breath hold: Maastrro experience

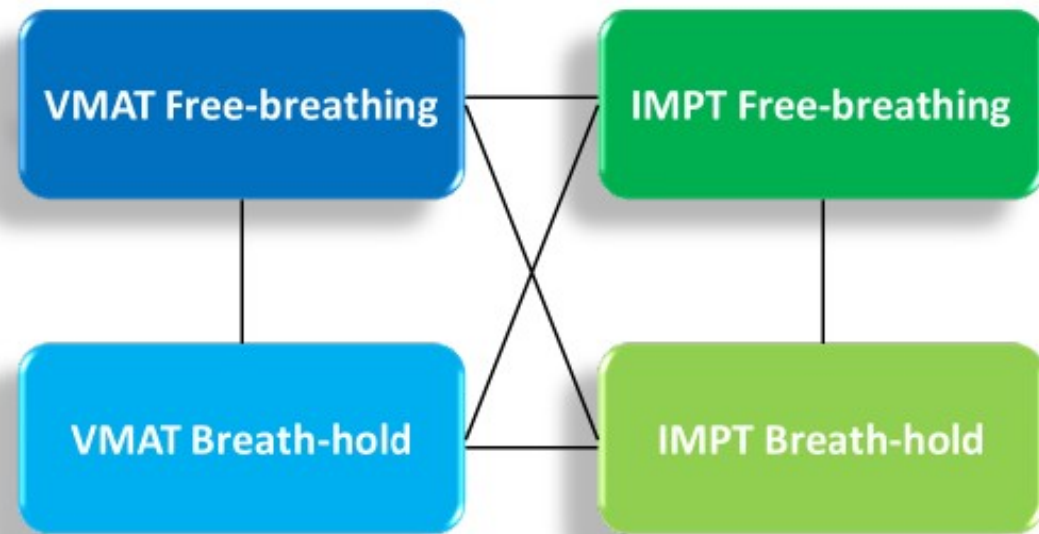


# Lymphoma, protons

## Background

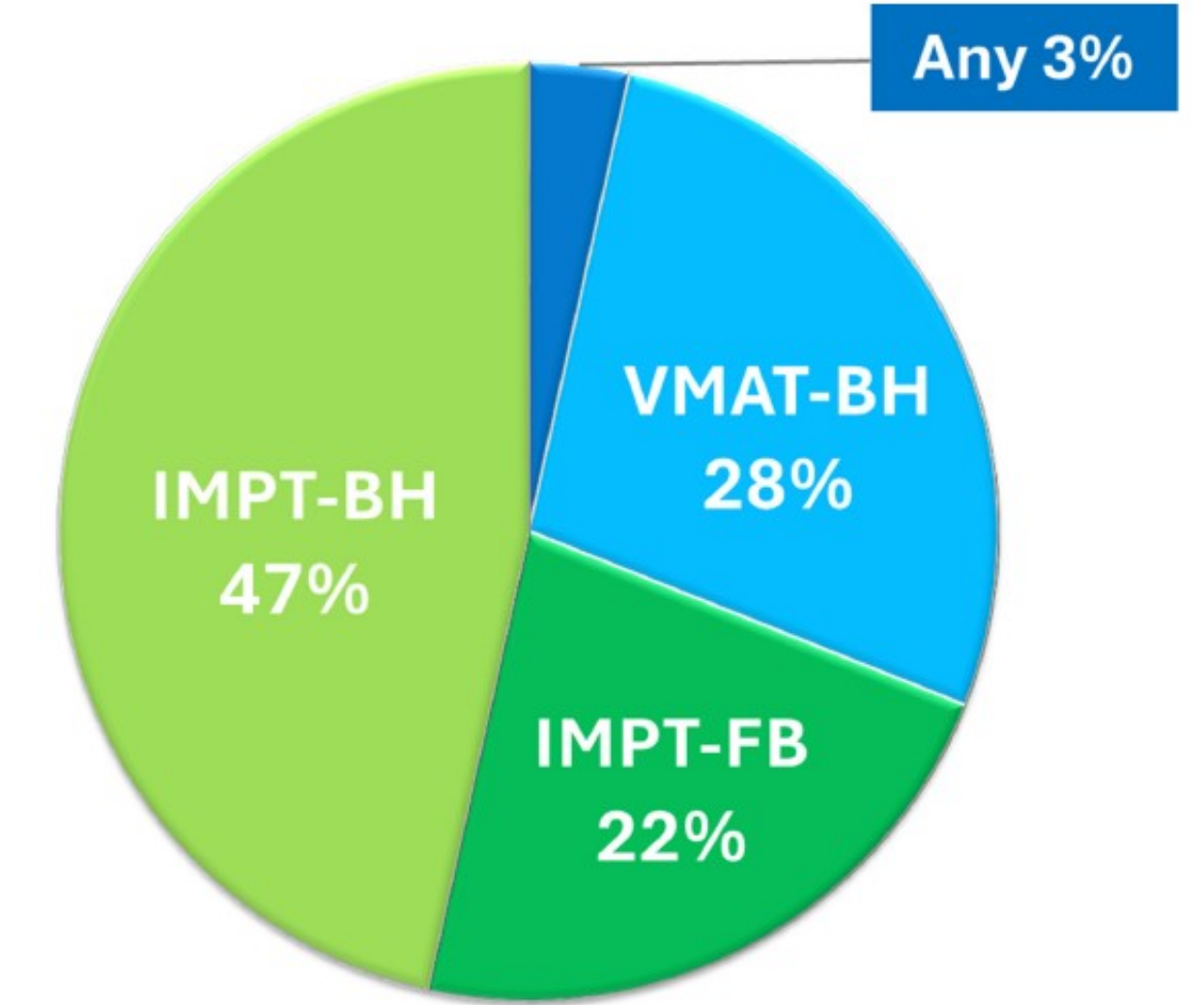
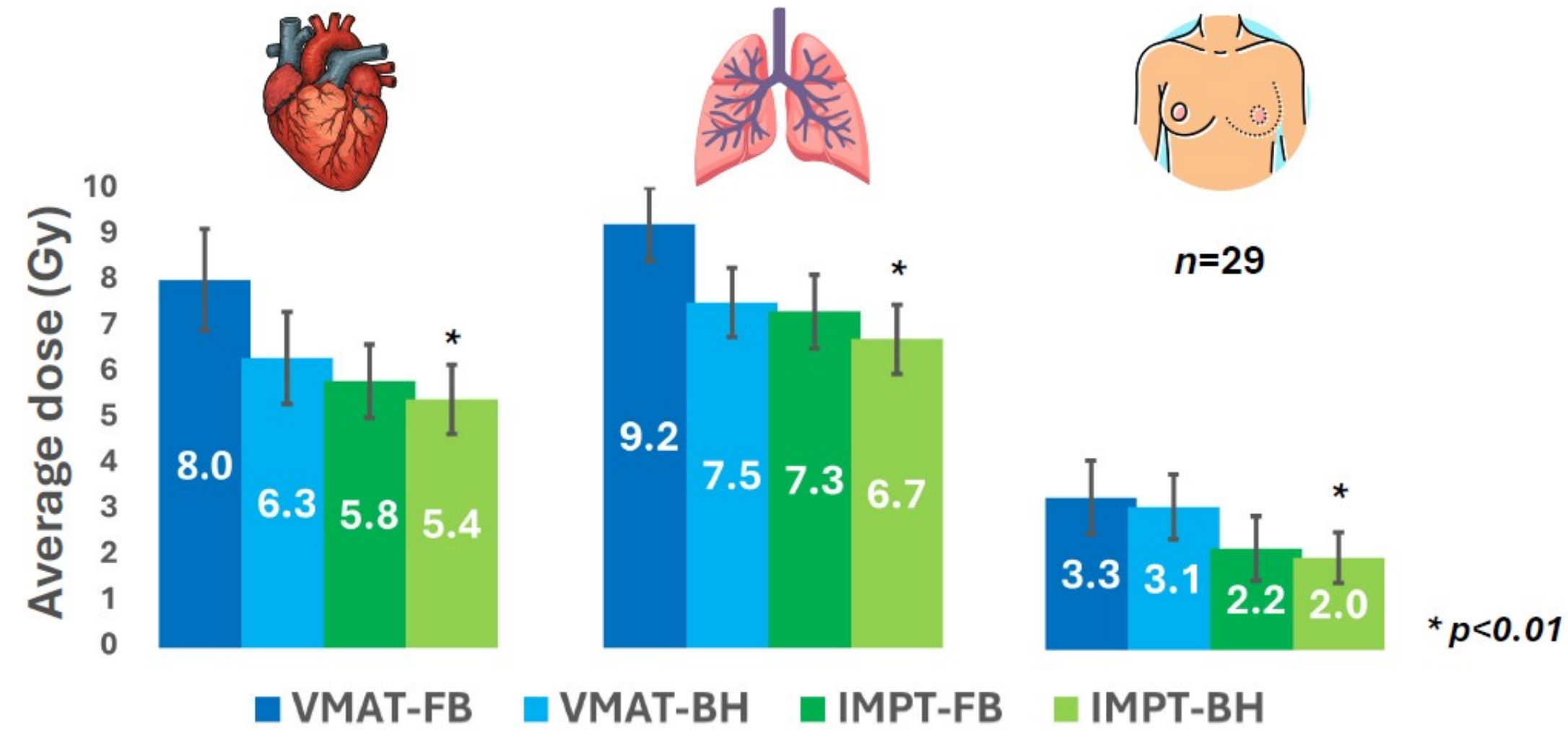
Comparative data is limited on **IMPT-BH** vs other techniques for **mediastinal lymphoma (ML)** patients.

## Material & Methods



- 58 ML patients
- 4 plans per patient: clinical + in silico
- DVH metrics

## Results



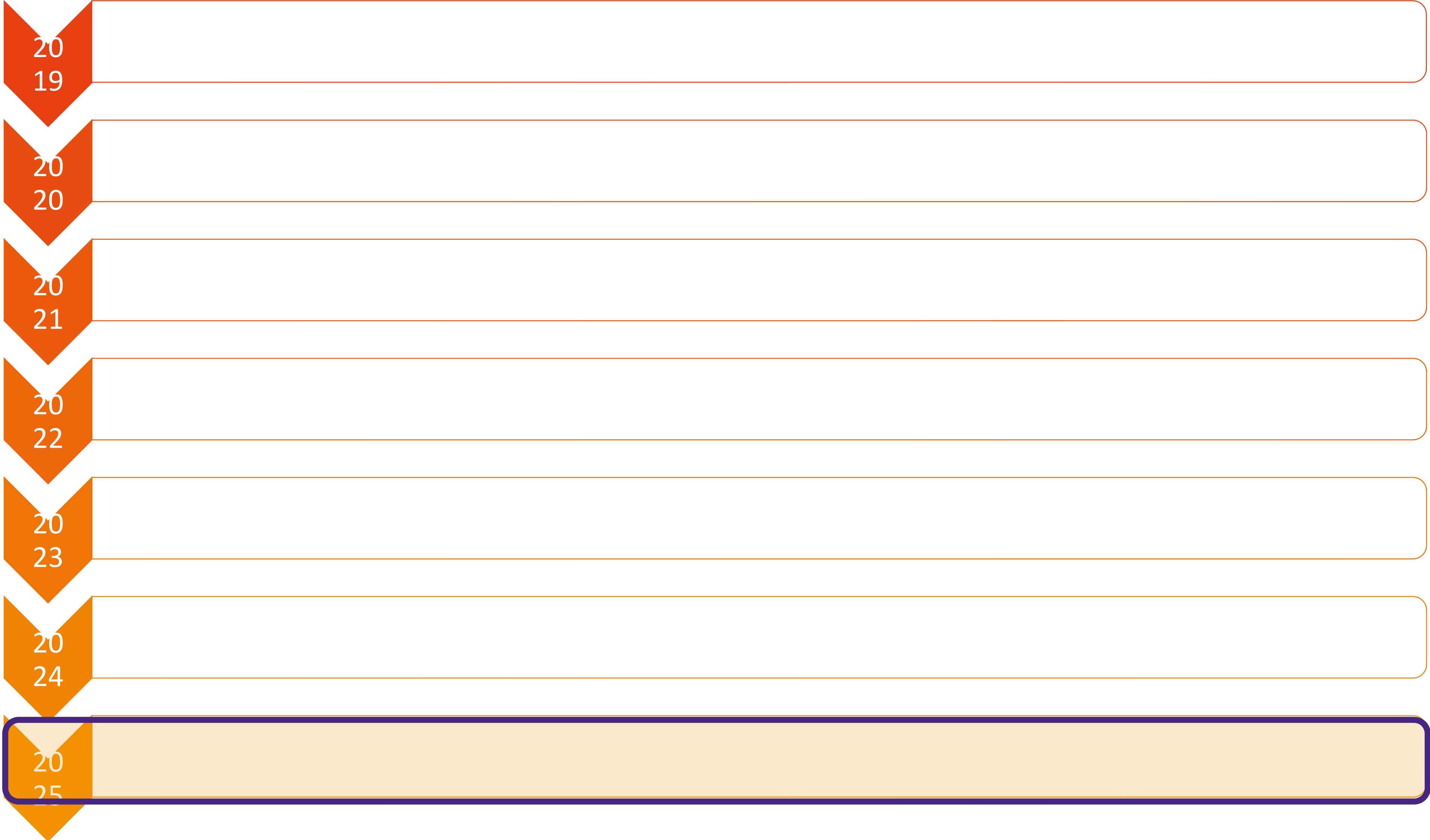
ΔDose ≥ 1 Gy = clinically relevant  
Patients assigned to overall optimal technique

## Conclusion

- IMPT-BH reduced mean heart and lung doses vs other techniques ( $p<0.01$ )
- IMPT reduced average bilateral mean breast dose vs VMAT ( $p<0.01$ )
- IMPT was the most optimal technique for  $\geq 69\%$  of ML patients

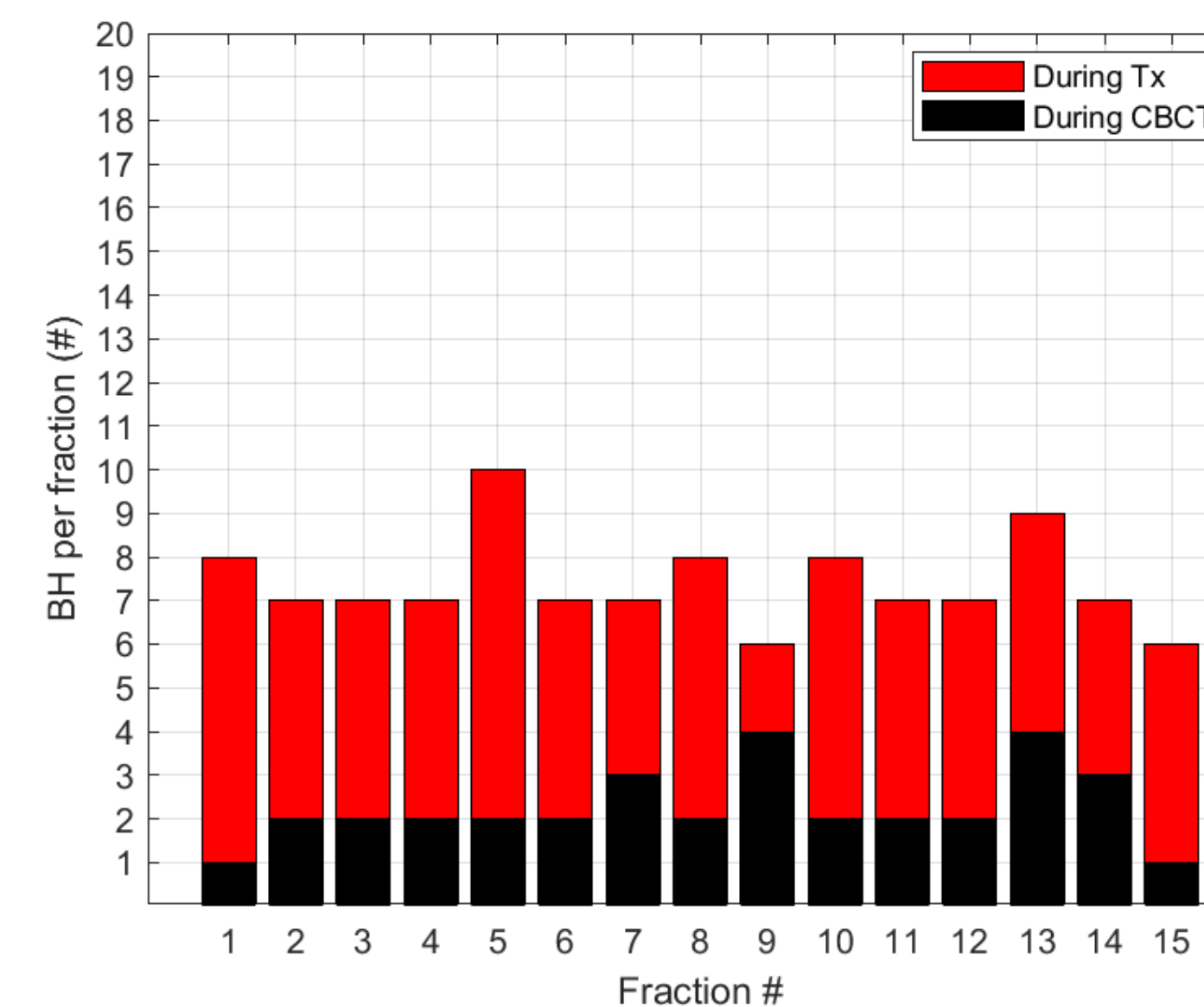
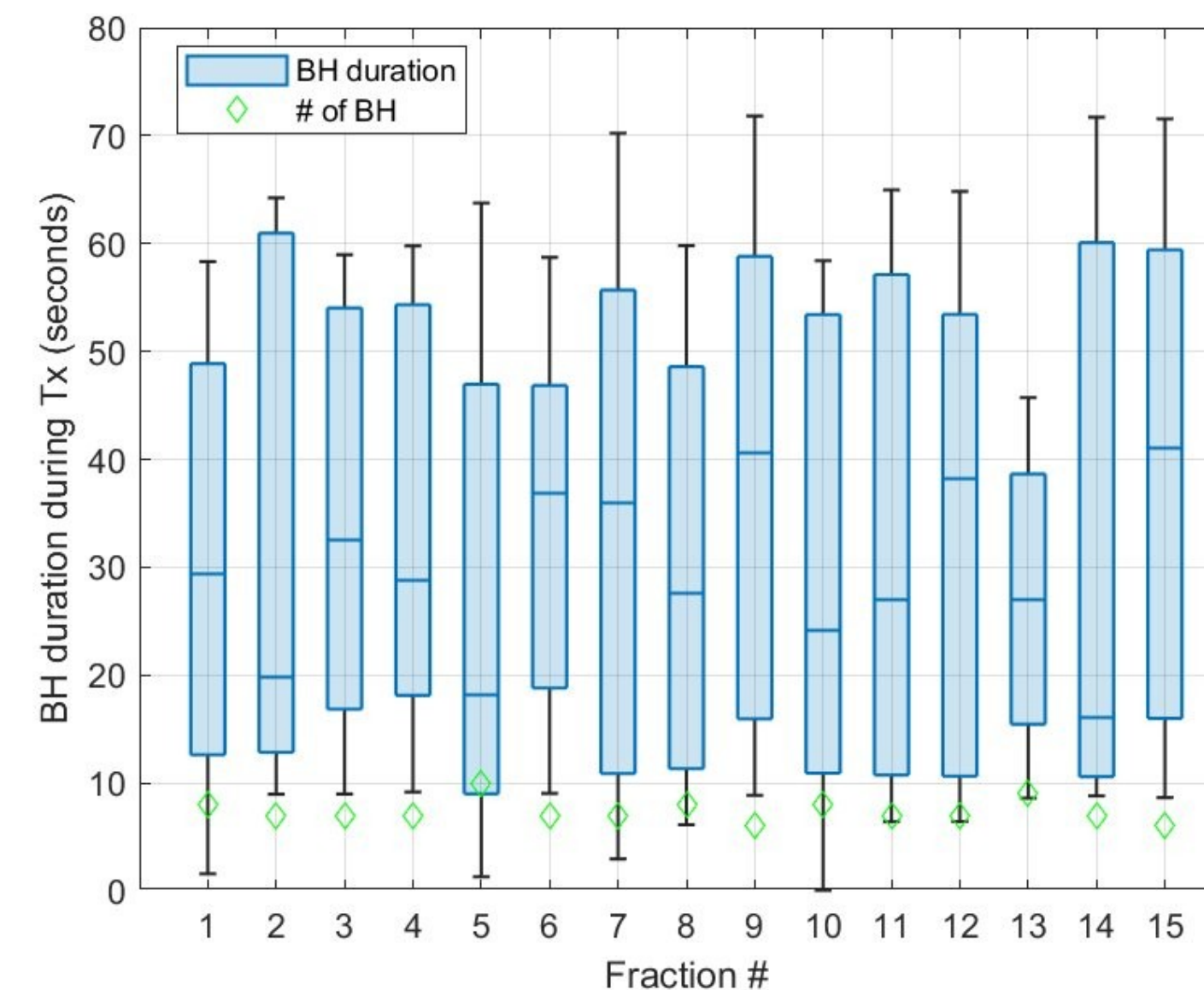
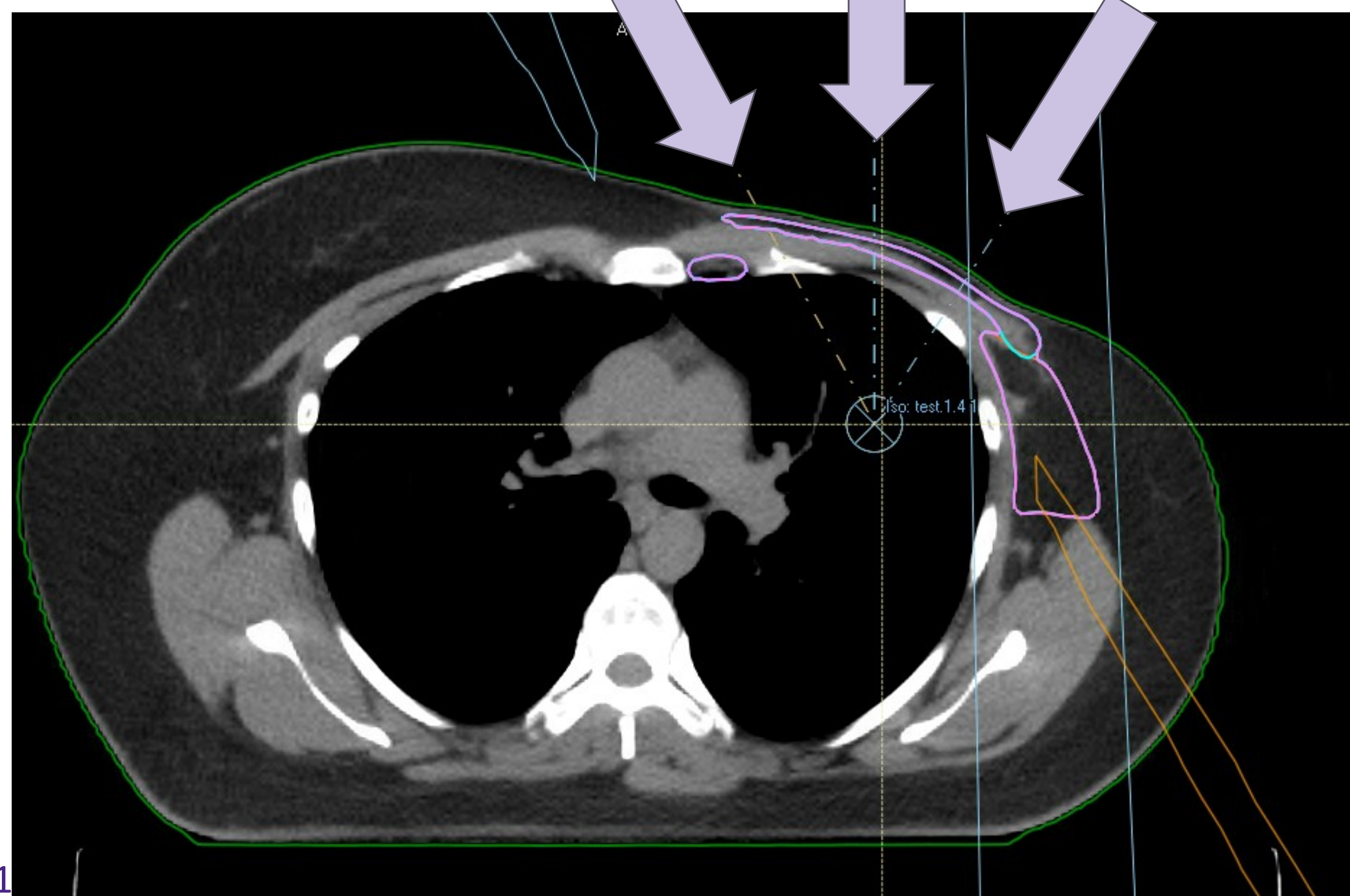


# NHFT in radiotherapy under breath hold: Maastrro experience



# Breast cancer, protons

- 1<sup>st</sup> patient (March 2025): 5 beams (3 directions), 15 fx



# Future work: ProThorac



Dose optimized positioning

UMCG:

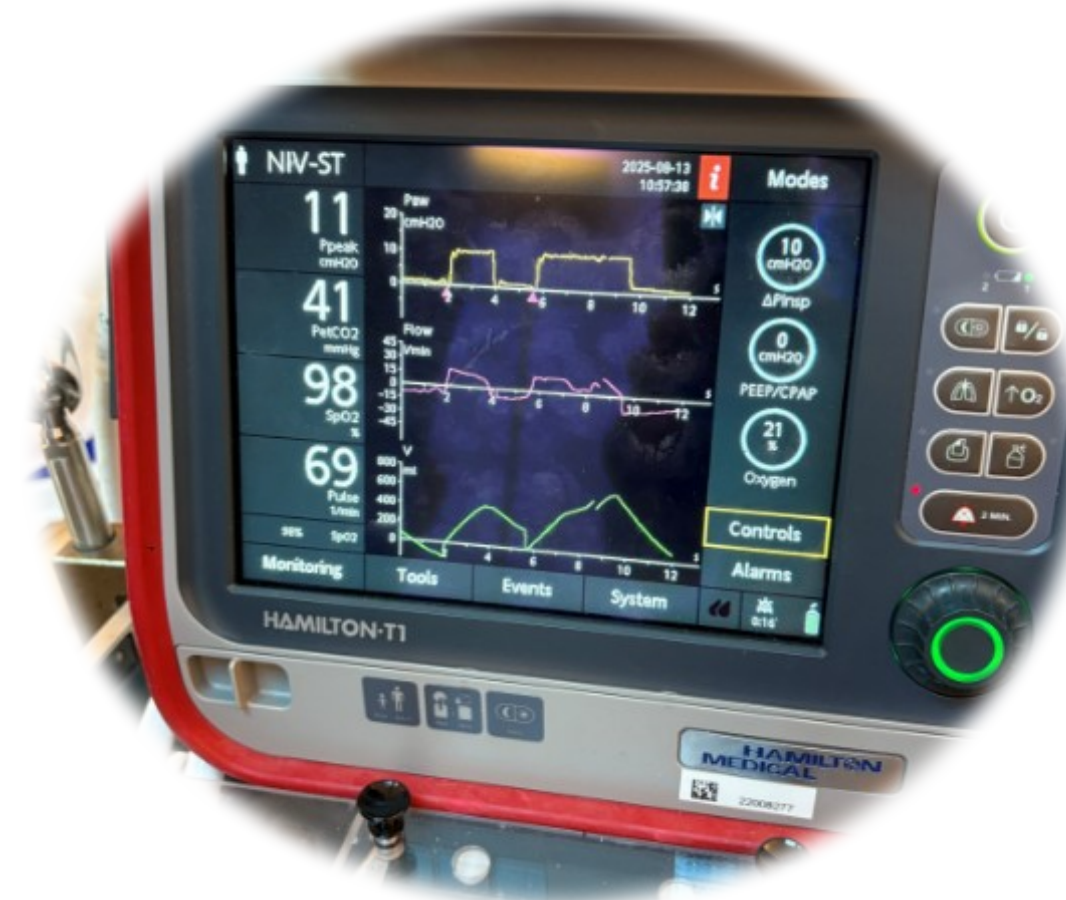


Probabilistic optimization

PROTONS  
OAR dose reduction in thoracic tumors

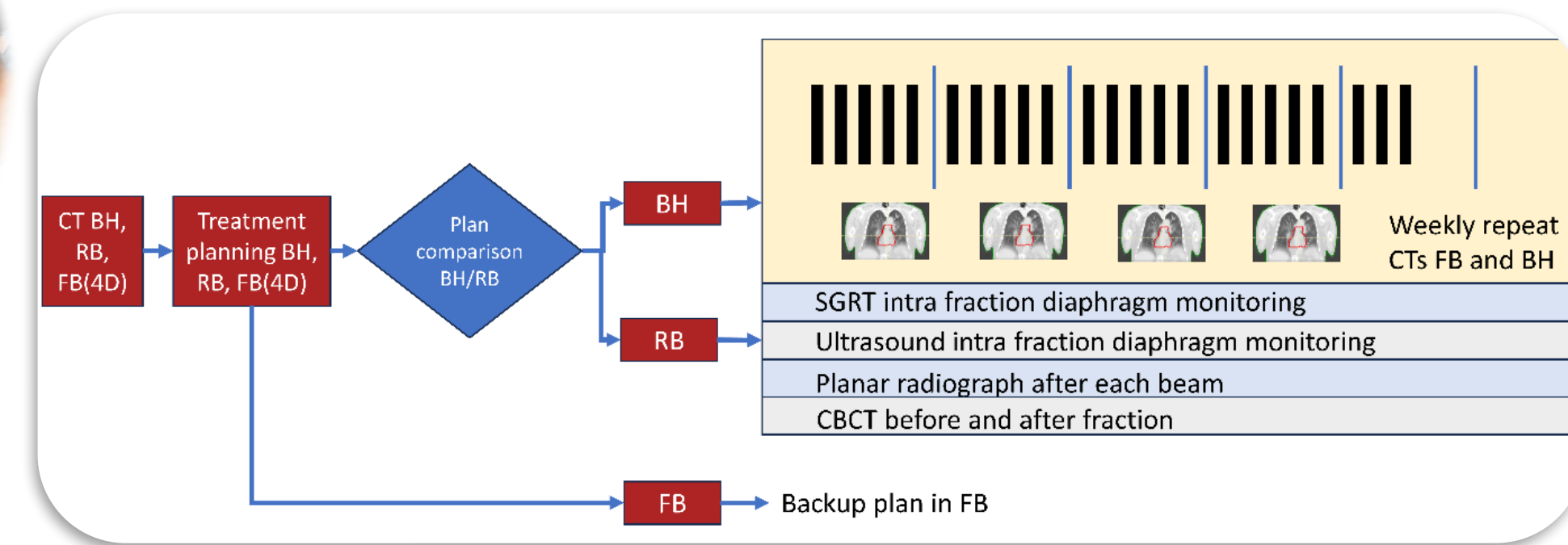
NIMV breath hold/regularized breathing

Maastrro



## Work packages

- In silico study to define most preferable tumor sites
- Develop a method for NIMV breath hold or regularized breathing
- Volunteer study
- Patient feasibility trial



# Teamwork

Thanks for slides to

Richard Canters

Bastiaan Ta

Gloria Vilches-Freixas

## RTTs

Kim van der Klugt

Fleur Vereijken

Maud Cobben

Maud van den Bosch

Indra Lubken

Anne van Engelen

Cissy Stultiens

Marije Velders

Femke Visser

Eva Rousch

Debby Tissen

Esther Van Enckevort

....

## RTOs

Bastiaan Ta

Maaïke Berbée

Karolien Verhoeven

Liesbeth Boersma

Stéphanie Peeters

....

Gloria Vilches-Freixas

Esther Kneepkens

Richard Canters

Giorgio Cartechini

Femke Vaassen

Colien Hazelaar

Ilaria Rinaldi

....

## Physicists

thank you