Welcome to Advanced Skills Modern Radiation Therapy

Prague 2017

Fourth run!

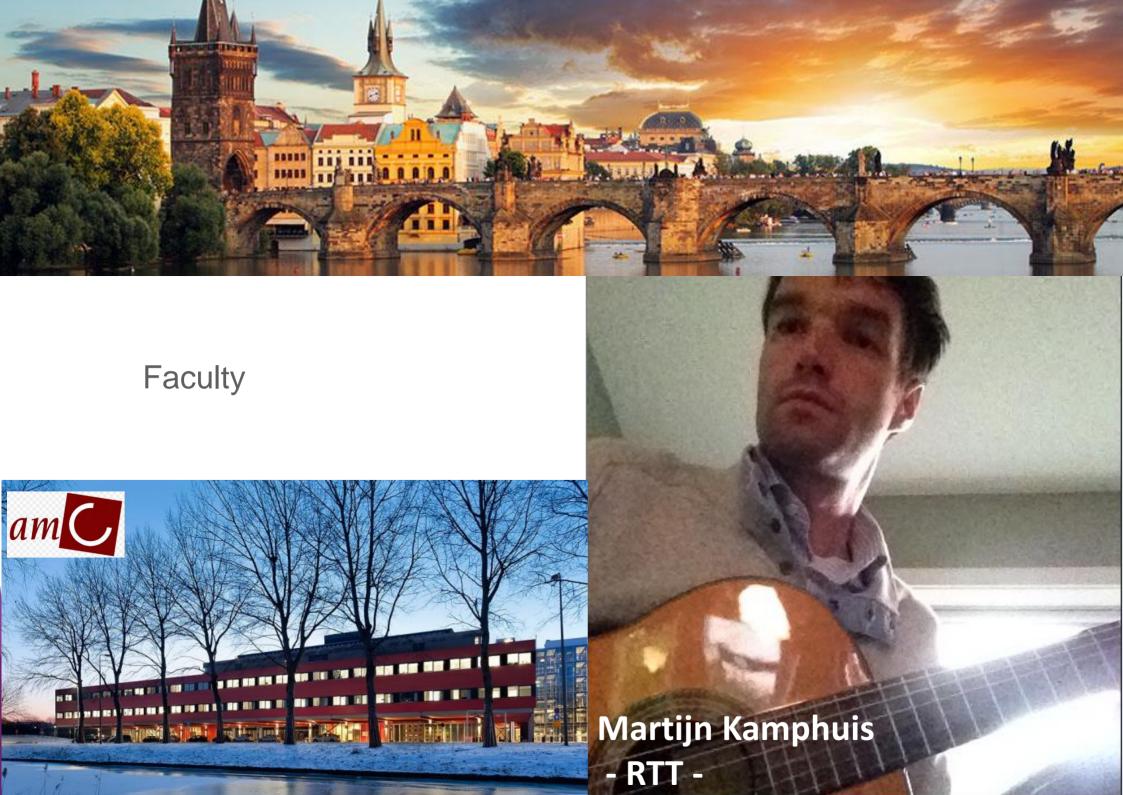
- Amsterdag
- Copenhagen
- o Dublin







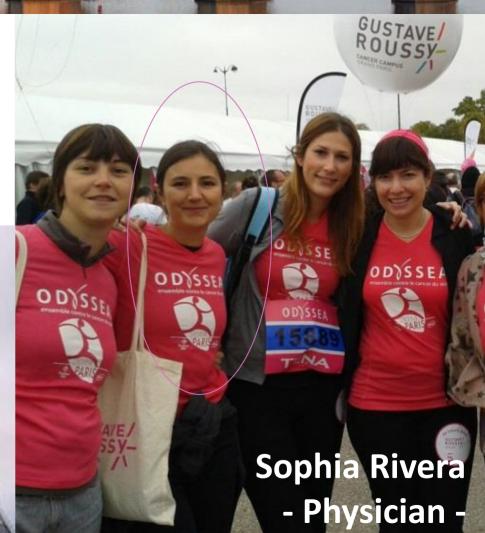


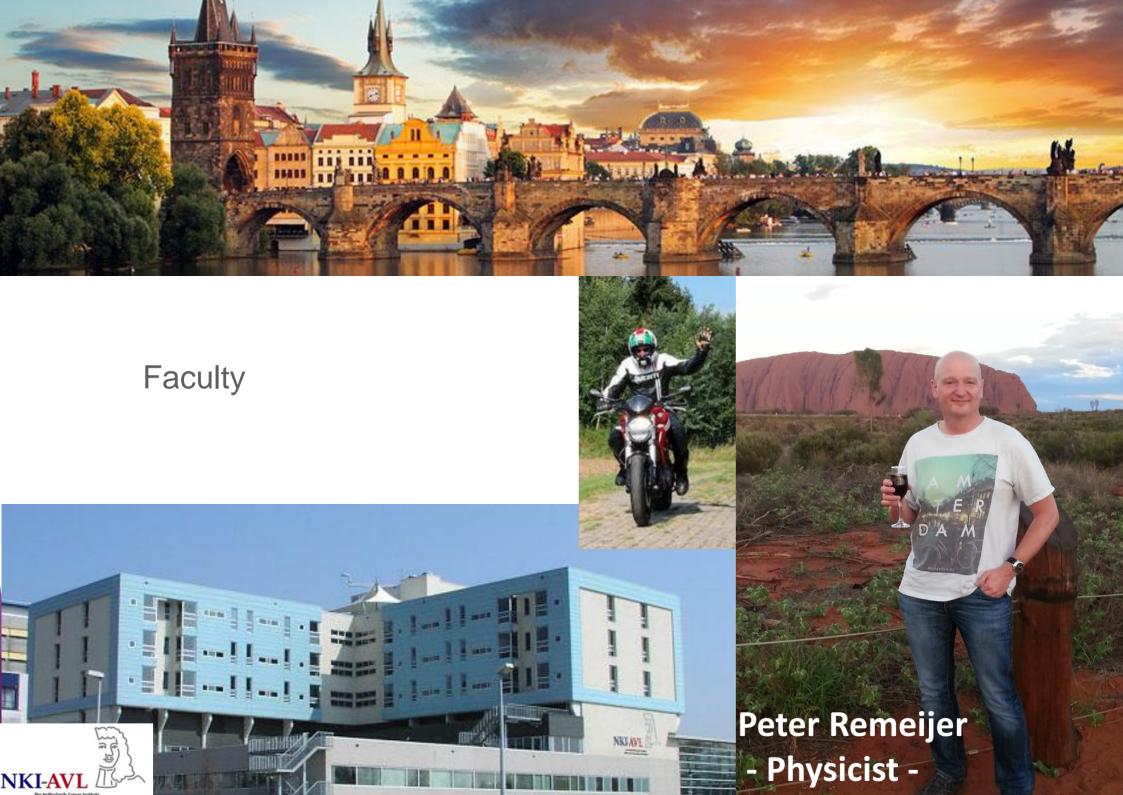


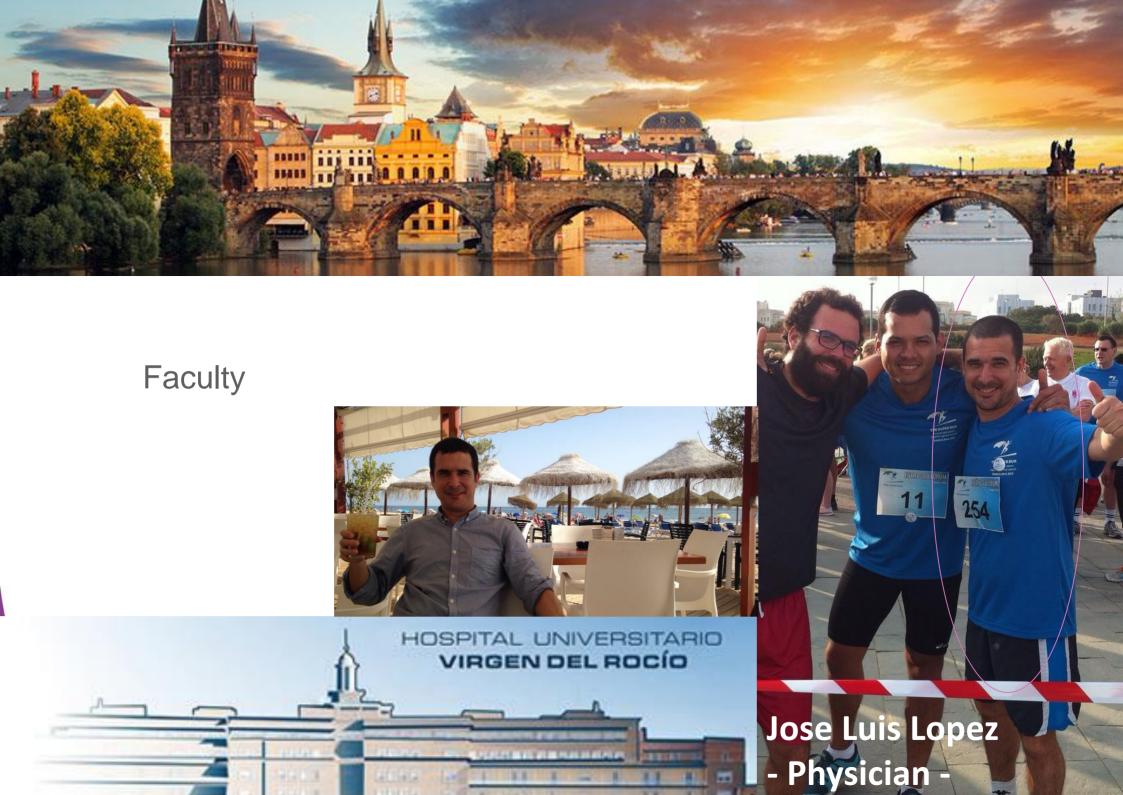


Faculty



























- 1 Australia
- 1 Austria
- 1 Bosnia Herzegovina
- 6 Czech Republik
- 5 Denmark
- 2 Malta
- 3 Poland
- 2 Portugal
- 2 Slovenia
- 1 Spain
- 2 Switzerland
- 5 Netherlands
- 3 Turkey
- 1 United Kingdom

34 Participants

+

7 Faculty

+

4 Company delegates

+

1 ESTRO









Program

4.5 days

24 lectures ~30 minutes

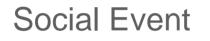
5 workshops

1 site visit

1 social event







Tuesday June 13

18.30 @Main Square Jan Hus Monument

Followed by Dinner







- All steps of modern Radiation Therapy -





Turning Point





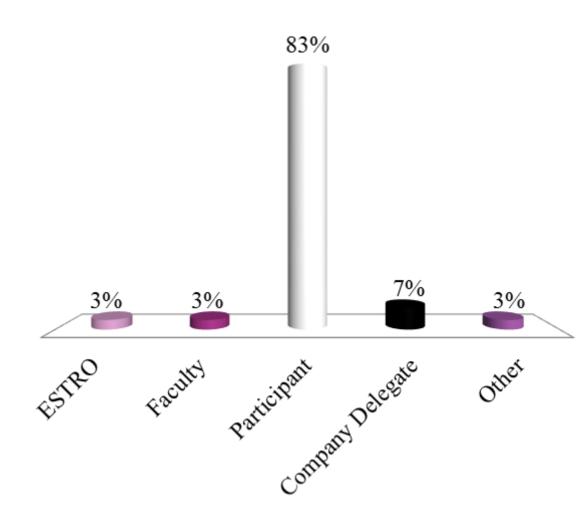
Turning Point

A little test!



Who is in the room?

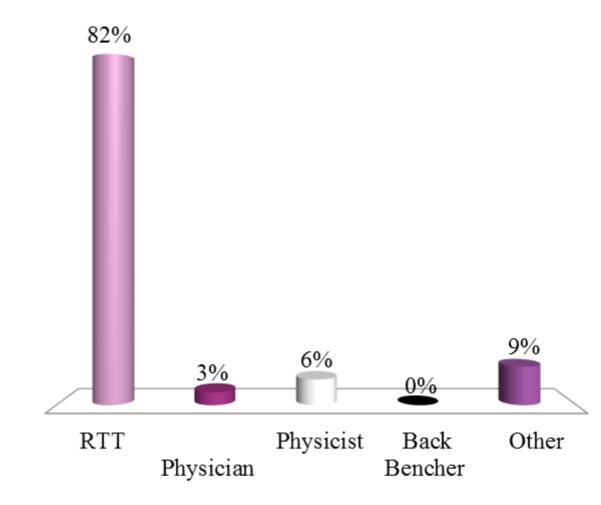
- A. ESTRO
- B. Faculty
- C. Participant
- D. Company Delegate
- E. Other





Who is in the room?

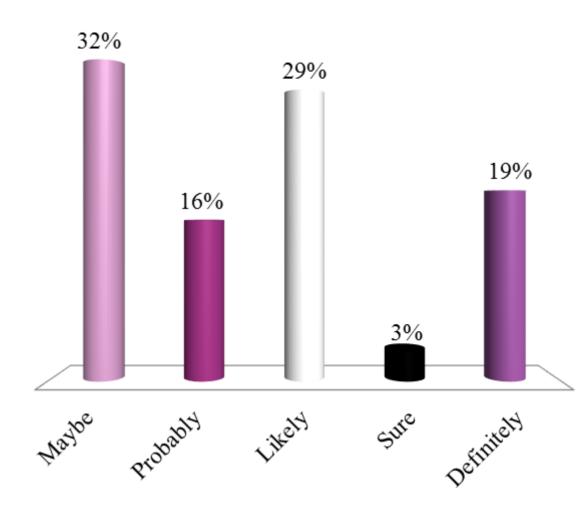
- A. RTT
- B. Physician
- C. Physicist
- D. Back Bencher
- E. Other





Who is joining for beers @end of day?









Laptops – workshops

- Delineation
- Margin calculation
- Safety issues & prospective risk analysis





Questions?







RTT's Perspective on modern Radiation Therapy

Rianne de Jong *RTT,*Academic Medical Centre
Amsterdam





Prague 2017
m.a.j.dejong@amc.uva.nl

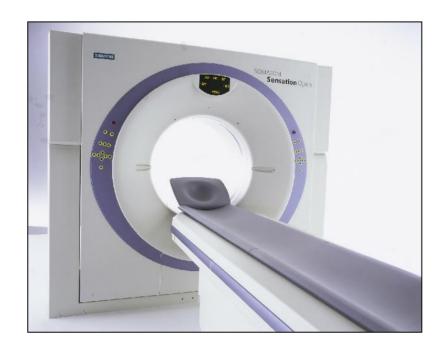


Changes over the last years

Simulation:

from fluoroscopy to CT



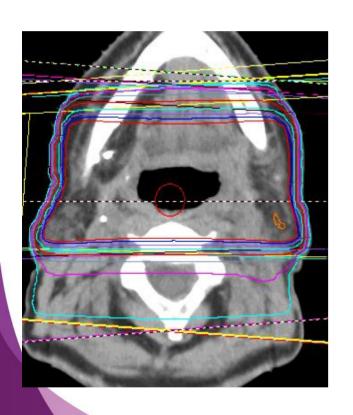


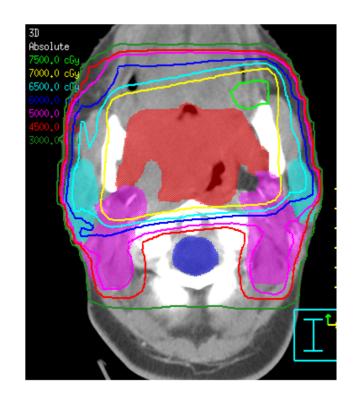
2 D 3 D

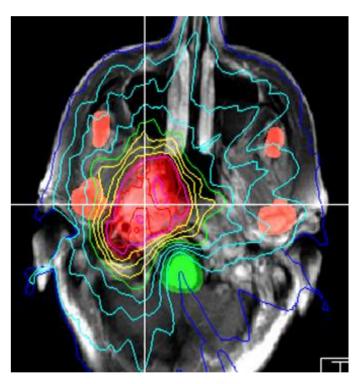


Treatment planning:

from conventional to conformal to IMRT & arc therapy









Treatment machine:

From patient set-up with skin marks to additional patient set-up verification

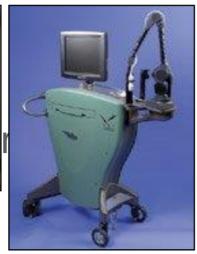
- Portal imaging (2D MV)
- Kilo voltage imaging (3D kV)

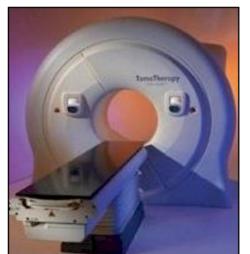




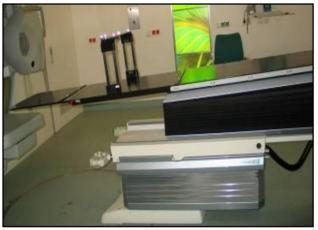










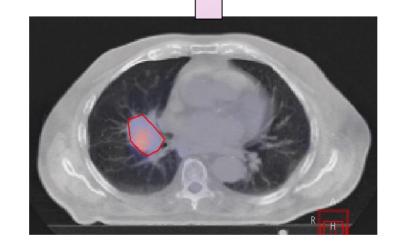




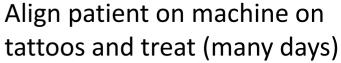
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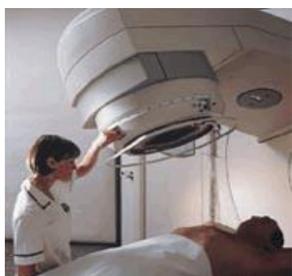
Tattoo, align and scan patient





Draw target and plan treatment on RTP



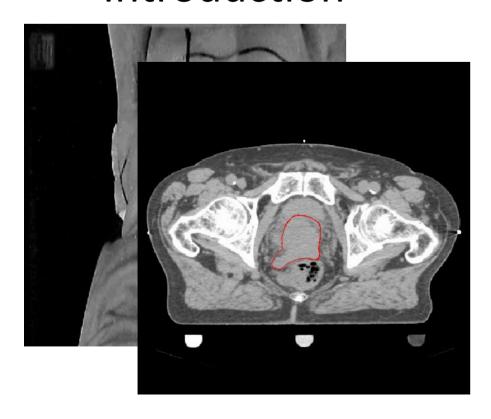


In principle this procedure should be accurate...

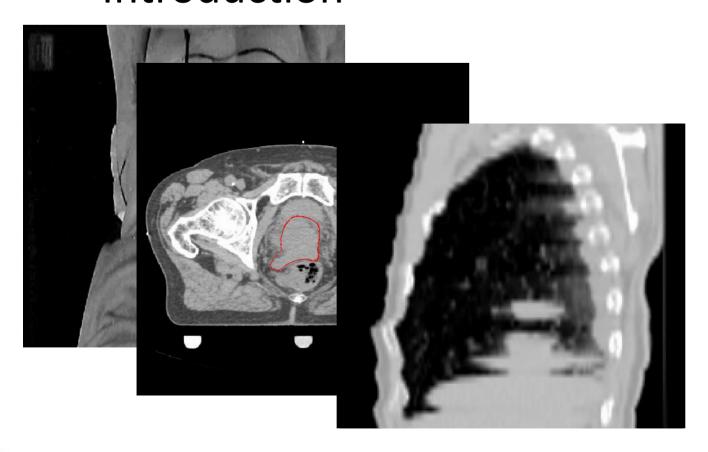




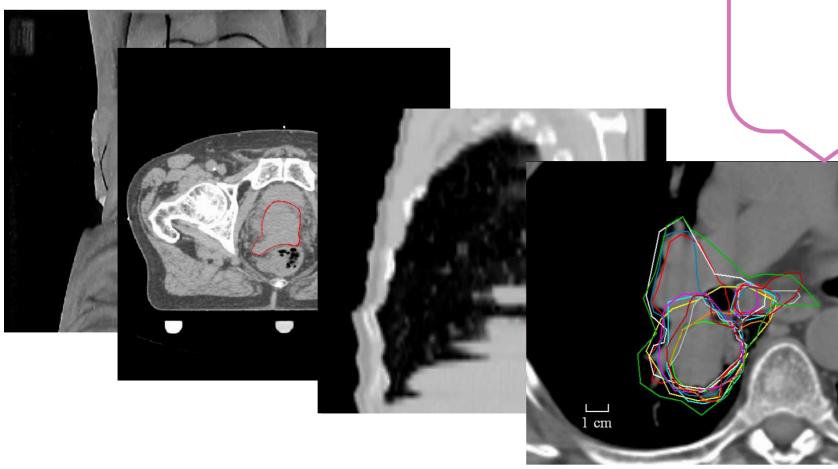






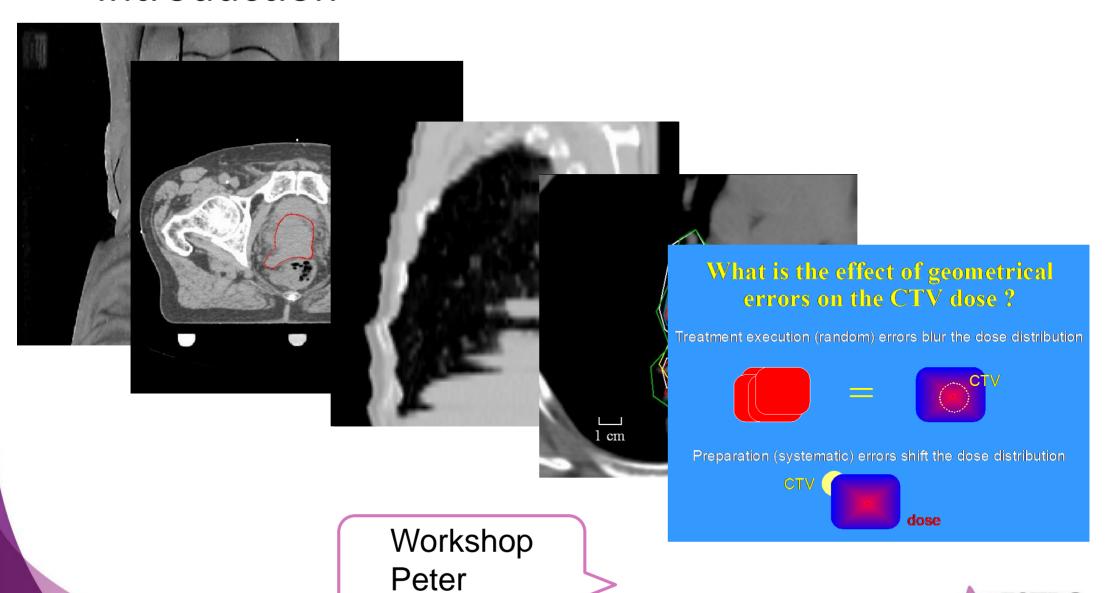






Workshop Sofia Elizabeth Jose Peter







PTT's Job







The RTTs job

- Patients education
 - Pre-treatment imaging
- Simulation
- Treatment Planning
- Treatment
 - Image guidance
 - Research & Development
 - Some sort of specialization in one step of the treatment chain: Sometimes controversial: all-round RTT is considered optimal job description.



Patient education

2 departments, 2 solutions:

AMC	Avl	_
/ \ \ \ \ \ \	/ \ V L	_

4 RTTs
 3 RTT's assistent

• 20% 80% time spent

• 30% 100% patient coverage

Combined not combined with working on treatment machines

Only 1 slide...? *Very important to the patient!*

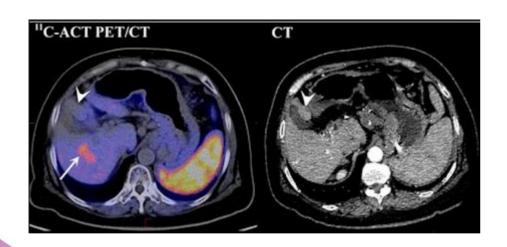


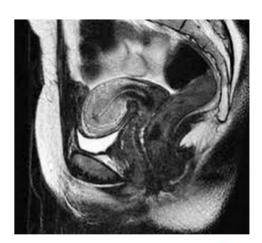
Pre-treatment Imaging: PET/MRI/CT

Often combined use with radiology department:

Always one RTT from radiation therapy

- Trained in delivering contrast agents
- Focused on patient positioning: registration images for delineation





Simulation CT

RTTs working on CT combined with working on the treatment machines Sub group only working on CT

- Contrast agents
- 4D CT
- Breath hold CT



Treatment Planning

RTTs working on Treatment Planning combined with working on the treatment machines.

Sub group working treatment planning only – research and development.

Physicist only in the loop when outside of tolerance Physician have to sign off on the plans

- Multi modality registrations
- Delineation of Organs at Risk
- IMRT VMAT (all curative intent treatments)



Treatment

3 RTTs per machine when breaks are scheduled

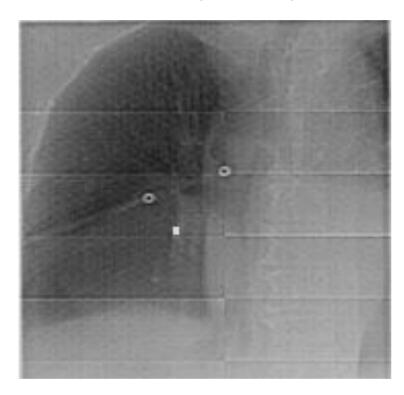
4 RTTs per machine for full program



Patient Support

Support patients and their relatives and friends:

During RT in RTT's working area for support and transparency



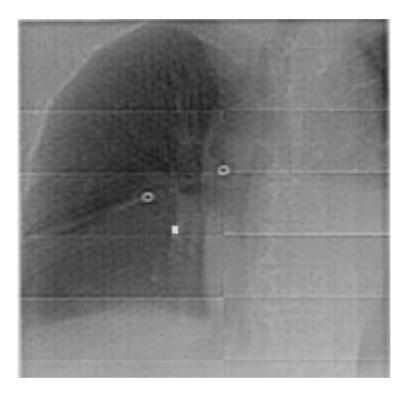
Portal image



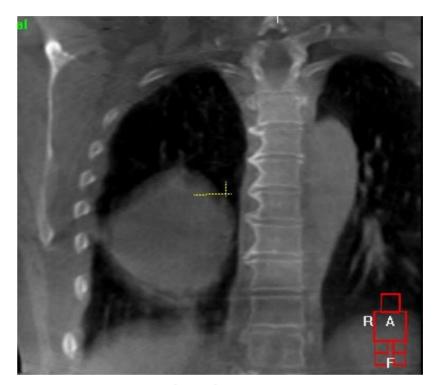
Patient Support

Support patients and their relatives and friends:

During RT in RTT's working area for support and transparency



Portal image



CBCT image



Starting IGRT (3d)



IGRT

- It is at the end of the treatment chain
- It involves all RTTs! Not only working on the treatment machine
- It requires understanding of all steps in radiation therapy
- It is still evolving: MRI-linac!



Implementing CBCT



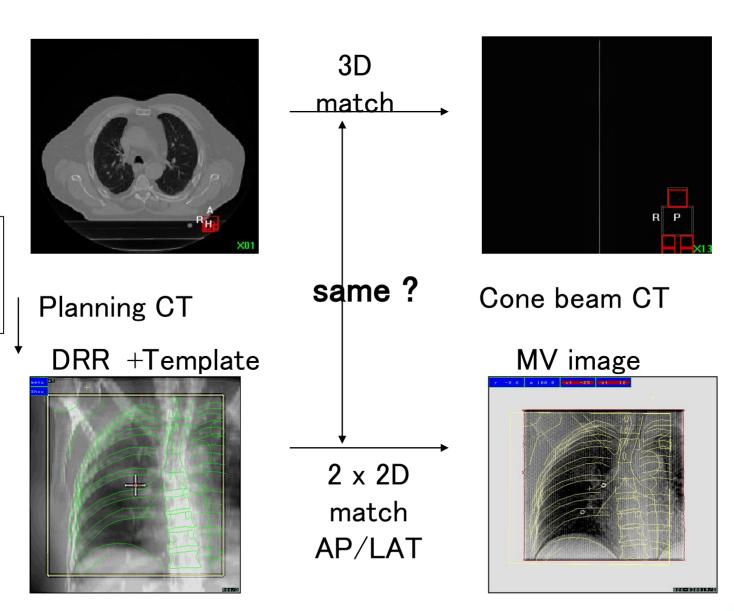
June 2003:

- 4 RTT's
- 2 Physicists
- Patient program in the morning
- CBCT in the afternoon
- 8 months of validation

Implementing CBCT: validation of the system

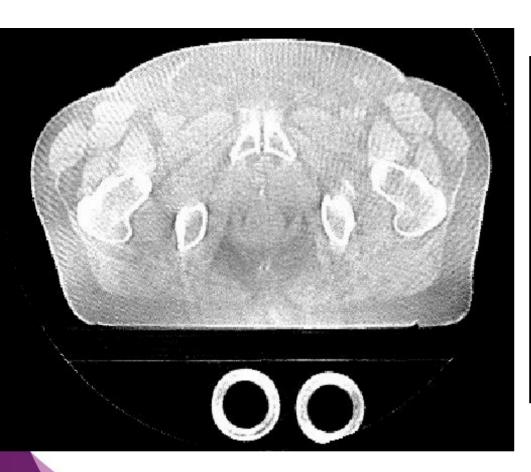
Cross

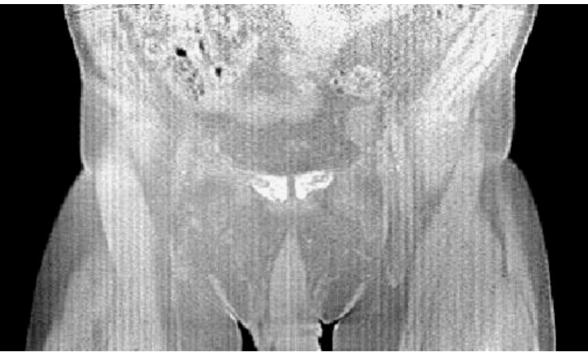
validation



Implementing CBCT: designing imaging presets

320 Projections 1.5 - 3 cGy

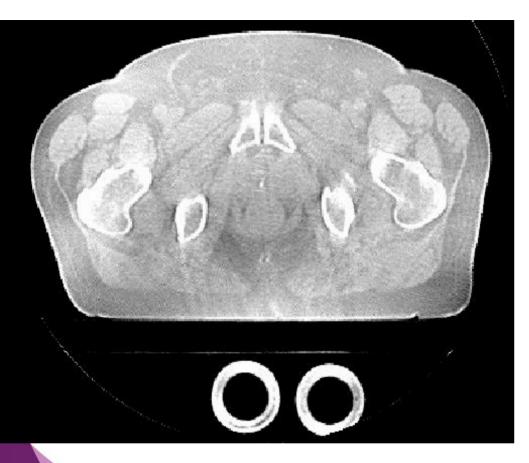






Implementing CBCT: validation of the system

640 Projections 1.5 - 3 cGy







Implementing CBCT: role of RTT

- Understanding basic physics and technical aspects of new imaging modality
 - IQ: artefacts
- Implementing in daily workflow
 - Protocols, manuals and working instructions
- Setting up training program for RTT's

Starting clinical use of CBCT

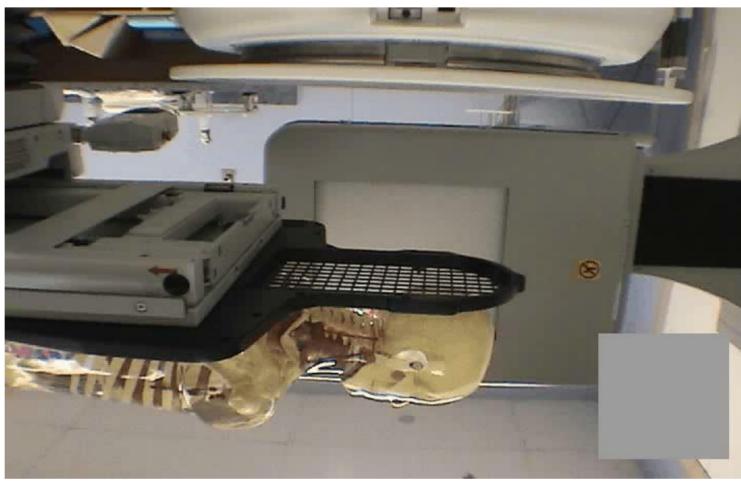
RTT's responsibilities:

- Acquisition of CBCT
- Registration bony anatomy (CBCT)
- Evaluation registration (CBCT)
- Evaluation of treatment!
- Execute decision rules off-line and online protocols

Same as portal imaging and a bit extra

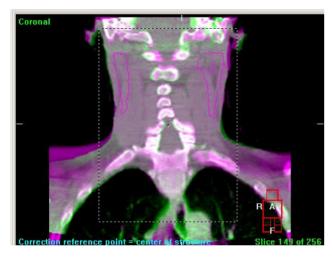
Clinical daily routine

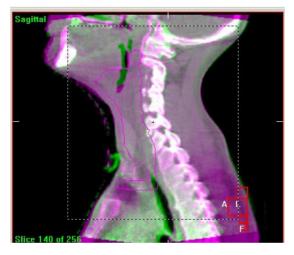


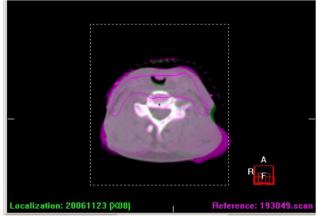


Courtesy to Doug Moseley (PMH) Jan-Jakob Sonke (AvL)

Clinical daily routine - registration

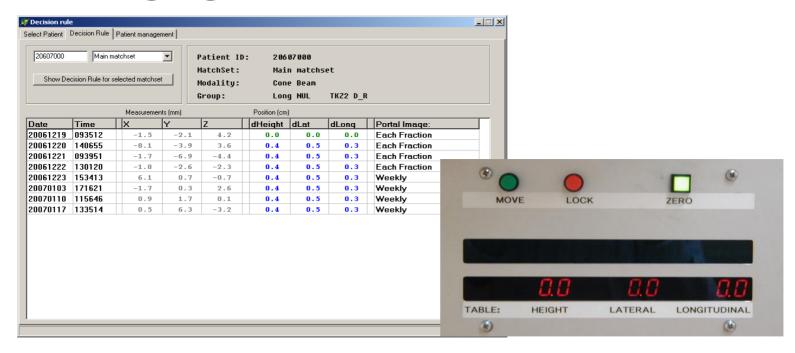


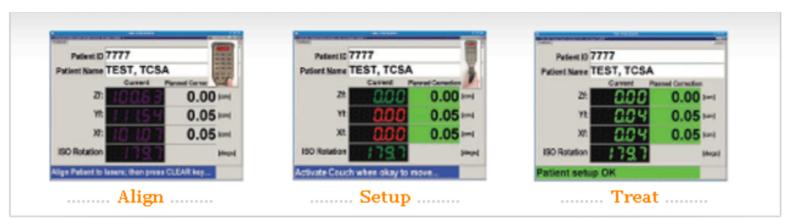




Automatic registration
CBCT scan

KV imaging – off/online correction





Managing IGRT (3d)



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5 RTT's with a focus on IGRT:

- Track, check patients
- First contact of changes occur
- Training and education
- Manuals and protocols
- Data collection & handling

Track & check patients

B PosVerQA 1.0						
File						
File Patient ID: 2193509 Course: 1 Mamma/Thoraxwand Herbestral Category: BREAST 174 Fraction Nr: 0 \$	Correct target in Theraview ? Correct beslissingsprotocol ? Juiste structuren ingetekend ? Correct clipbox ? Correct correction reference point Parallel toestel ingevoerd ?	n.v.t. 🗸	PosVerQA 1.0 File Patient ID: 2193509 Course: 1 Mamma/Thoraxwand Herbestral Category: BREAST 174 Fraction Nr: 1 \$\frac{1}{2}\$	Eenmalige check Wekelijkse follow-up Alle Items correct afgevinkt? Nieuwe set-up correctie juist overgenomen? CVT binnen PTV? Veranderde pathologie? Maximale afname in bodycontour? Blaasvulling voldoende? Afgevinkt? Rotatie binnen protocol? Ligt patient vergelijkbaar op CBCT als op CT? Welk plan is geselecteerd? Was er een tweak nodig?	Anatomische verandering Anatomische verandering Rotaties (>-4) Positioneringshulpmiddel ART Markers	vrije tekst
				Waren alle markers nog aanwezig ? Heeft er migratie plaatsgevonden ?		•
					Modify Remove	



@AMC

5 RTT's with a focus on IGRT:

- Track, check patients
- First contact of changes occur
- Training and education
- Manuals and protocols
- Data collection

Anatomical Changes

RTT should be trained in:

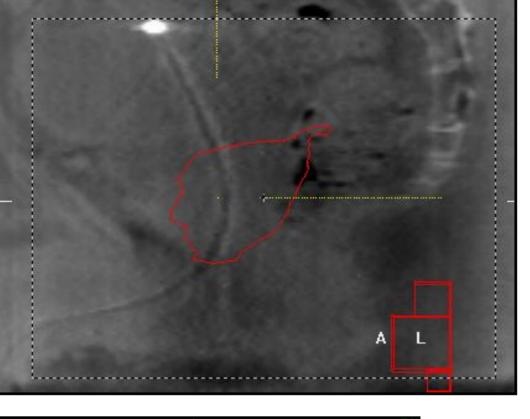
Recognizing patient changes/anatomical changes that have an influence on radiation treatment: Target coverage and/or dose distribution



RTT should have:

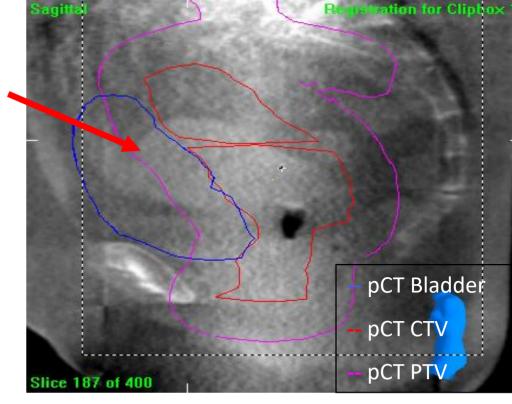
a management system for anatomical changes that flag the changes that may need intervention of some sort.

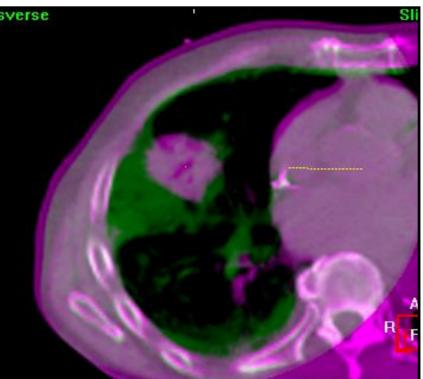


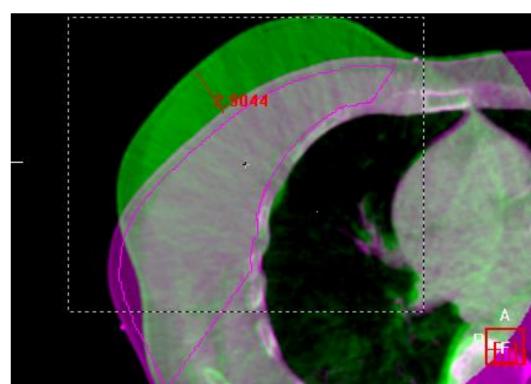


Ref CT

CBCT







Anatomical Changes

The important questions:

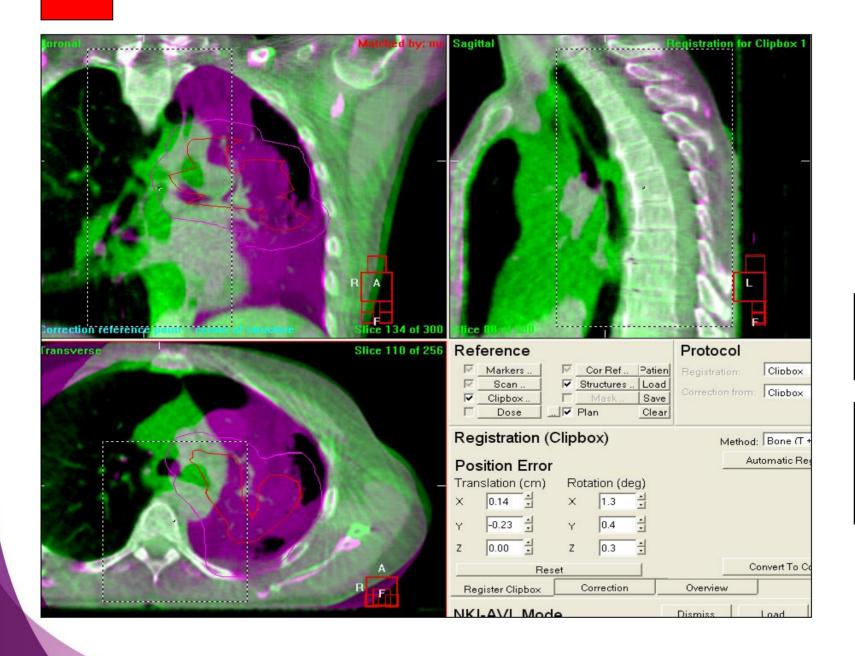
- 1: Is the target volume (CTV or GTV) within PTV?
- 2: Is the dose distribution compromised?
- Level green, no action needed.



- Level yellow, the radiation oncologist is notified by email, but no response is required to continue treatment.
- Level orange, the treating radiation oncologist (or back-up colleague) is informed by email and a response is required before the next fraction.
- Level red changes, the radiation oncologist must be consulted immediately before the treatment fraction is allowed to be delivered.



Level 1 Atelectasis resolved



GTV is not within PTV

Dose distribution is compromised



Anatomical Changes

Or keep it very simple:

Contact the IGRT-group when

- GTV is outside of PTV
- Anatomical changes > 1 cm



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5 RTT's with a focus on IGRT:

- Track, check patients
- First contact of changes occur
- Training and education
- Manuals and protocols
- Data collection

3 lectures (1h)

- Theraview: Portal imaging system and decision rule management system
- geometrical errors & correction strategies
- CBCT incl artefacts, image quality
- 2 Workshop (2x1.5h) in registration and image evaluation

Challenge: it affects all RTT's, so large group needs to be trained and kept up to date!



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5 RTT's with a focus on IGRT:

- Track, check patients
- First contact of changes occur
- Training and education
- Manuals and protocols
- Data collection



5 RTT's:

- Track, check patients
- First contact of changes occur
- Training and education
- Manuals and protocols
- Data collection

These RTT's also work in the clinic



Implementing IG&ART

Research department Clinic

Multi disciplinary group to implement, research and evaluate IGRT protocols:

- Physicists
- Physicians
- RTT's
- Software developers
- Post-docs/PhD students



Introducing IGRT

Also applicable for development steps of the radiation in other chain!

Also applicable for development of the radiation in other

RTT:

Evaluation of bulk of data: for example

- Inter fraction set up variability
- Intra fraction stability
- Organ motion or deformation
- Testing new (software) tools

Design & implementation new protocols

Training and education in house

Protocols and manuals

Clinic!

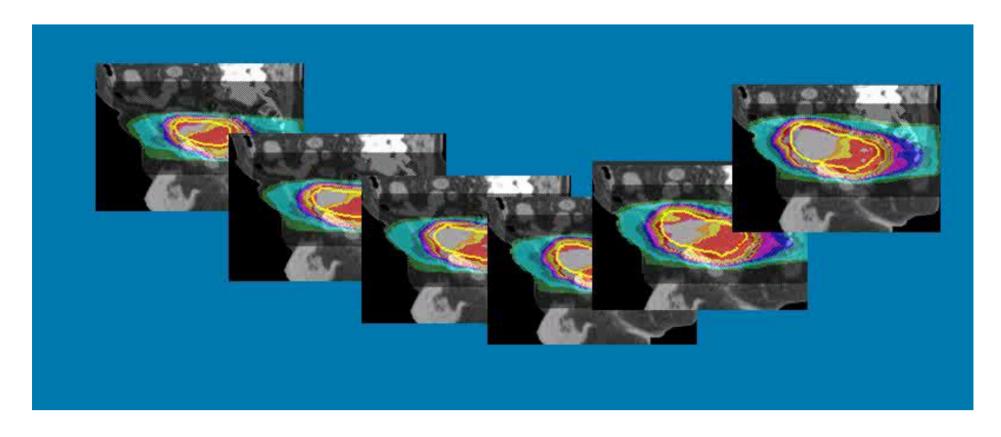


Shifting responsibilities @ treatment machine



ART: Library of Plan

Dealing with daily volume changes



Courtesy Danny Schuring, Catharina Ziekenhuis, Einhoven



Treatment Procedure

- Lipiodol demarcation of tumor by urologist
- Full & empty bladder CT scan
- Instructions to ensure full bladder
 - Good hydration prior to treatment
 - Empty bladder 1 hr before treatment
 - Drink 2 3 glasses
 - Continuous steering during treatment
- Cone-beam CT at start of treatment
- Selection of "plan of the day" based on bladder filling



Courtesy Danny Schuring, Catharina Ziekenhuis, Einhoven



Daily plan selection

Daily plan selection at linac



Shift in responsibilities!

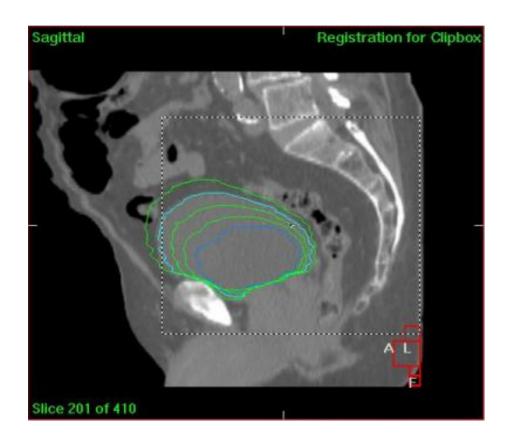




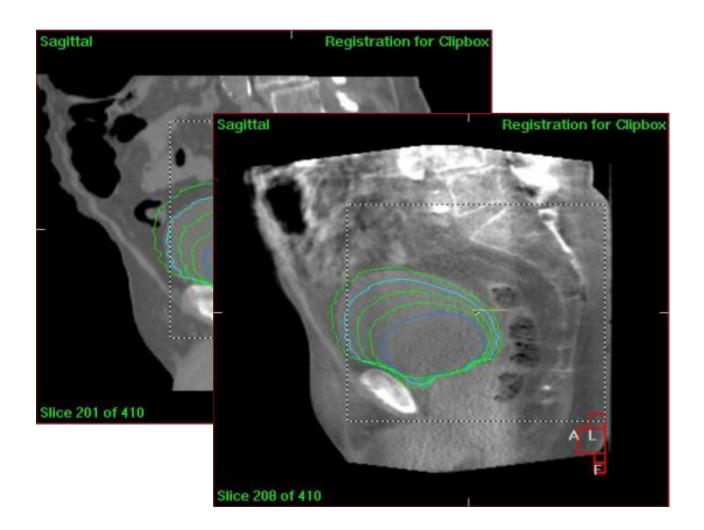
• Current practice: selection by physicist or specialized technologist



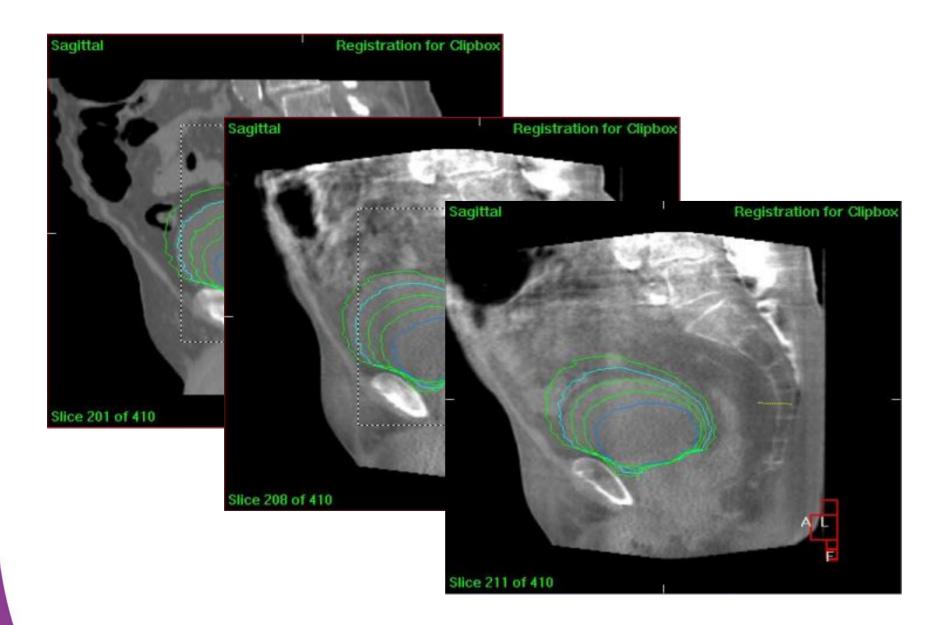








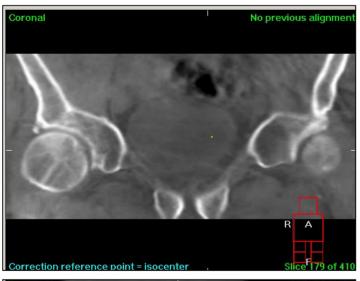




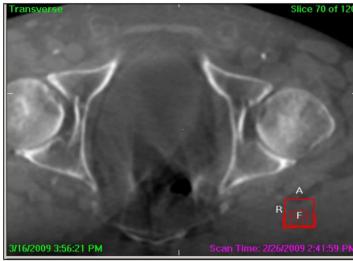


XVI quality

Workshop Rianne

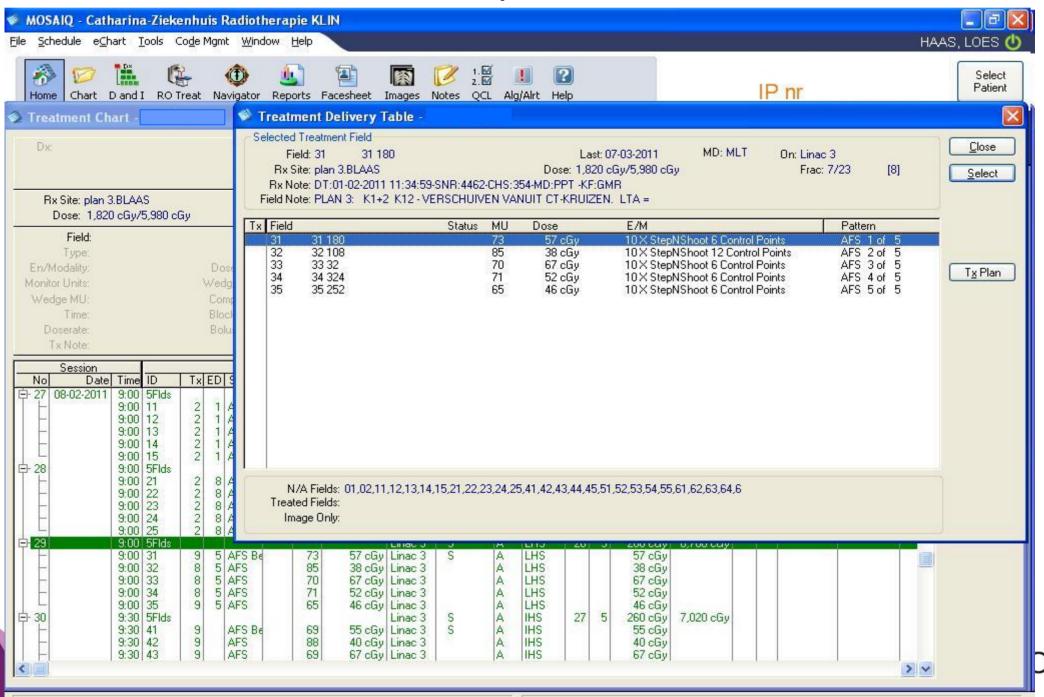




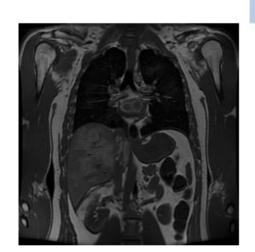




Plan selection in Mosaiq

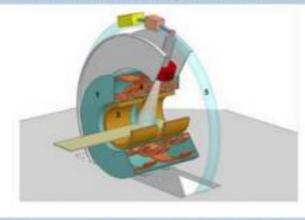


1 step further; MR inside the treatment room





U Utrecht-Philips Elekta



Viewray-MRIdian MRI-Cobalt



http://www.viewray.com

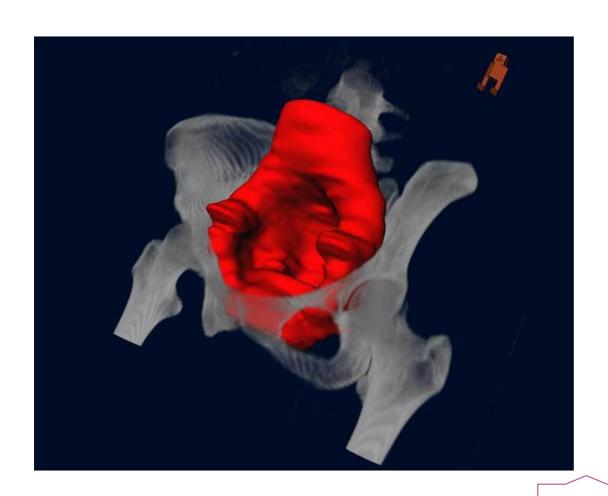
Diagnostic quality scan at treatment

Allows for:

- online re-planning
- online correction intrafraction motion
- ART: accumulate doses for adaptation
- Treatment response assessment for adaptation



MR for online replanning – needs contouring



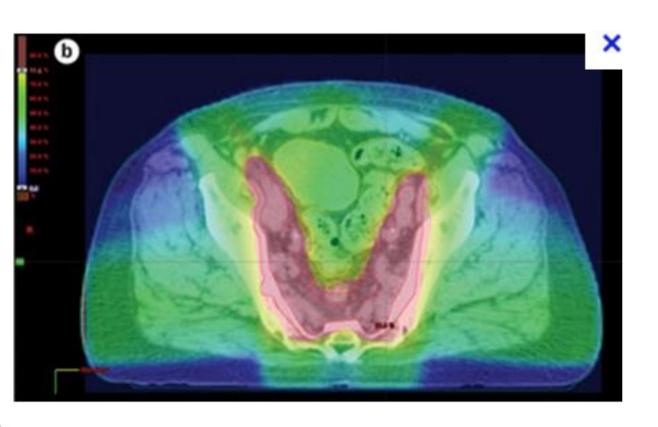
Approval of segmentation?

- OAR's
- Target volume

Peter



MR for online replanning – needs replanning

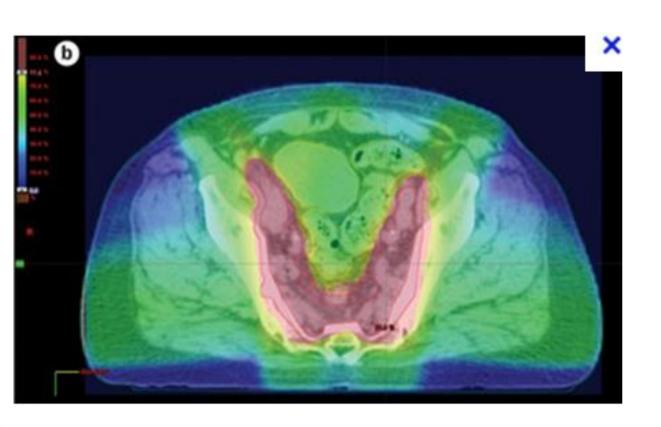


Approval of new plan?

- OAR's
- Target volume



MR for online replanning – needs replanning



Approval of new plan?

- OAR's
- Target volume

Treatment planning & IGRT become best friends!





Summary

Modern Radiation Therapy is a multi disciplinary effort Modern Radiation Therapy has openened up the field for RTTs:

- Patients education
 - Pre-treatment imaging PET/MRI/CT
- CT simulation
- Treatment Planning
 - Research and Development
- Treatment
 - Image guidance
- Research & Development



Acknowledgments

AMC

Coen Rasch
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AvL/NKI

Marcel van Herk
Peter Remeijer
Jan-Jakob Sonke
Anja Betgen
Suzanne van Beek

Catharina Ziekenhuis

Danny Schuring



Questions & Discussion



m.a.j.dejong@amc.uva.nl





Patient Preparation and Positioning

Martijn Kamphuis MSc MBA

(Slides: Rianne de Jong) Academic Medical Center, Amsterdam Prague 2017





Aim of Patient preparation and positioning

- → Minimize the difference in patient position
 - 1. between simulation and treatment sessions
 - 2. during the treatment session
- Maximize the distance between target volume and organs at risk

Tools:

- Immobilization and fixation
- Patient compliance

Tools of Patient preparation and positioning

→ Immobilization

Daily set-up **reproducibility** and **stability** through the use of fixation or aiding devices











Expectation management

- This aim of this talk is not to show the best devices
- Understanding the rationale behind it
- Choice for device will be based on:
 - **Economics**
 - Local availability
 - Literature
 - > Experience
- Link to important review at the end of the .ppt



Tools of Patient preparation and positioning



"My diabetic research shows that test subjects are 98% more likely to take their diabetic pills if the pills are covered in chocolate."



Minimize the difference in patient position

- Minimize the difference in patient position
 - 1. between simulation and treatment sessions
 - 2. during the treatment session
- Maximize the distance between target volume and organs at risk

Tools:

- Patient compliance
- Immobilization and fixation

Aim of Patient preparation and positioning

Minimize the difference in patient position between simulation and treatment sessions: *inter*-fraction motion

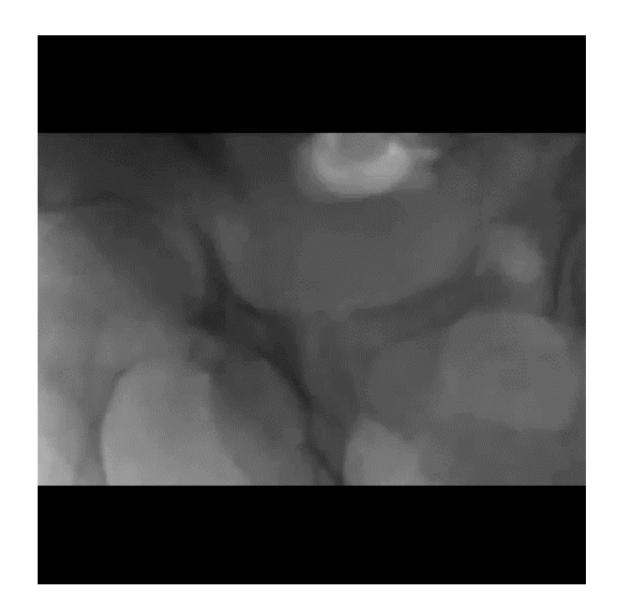
Tools:

Patient compliance:

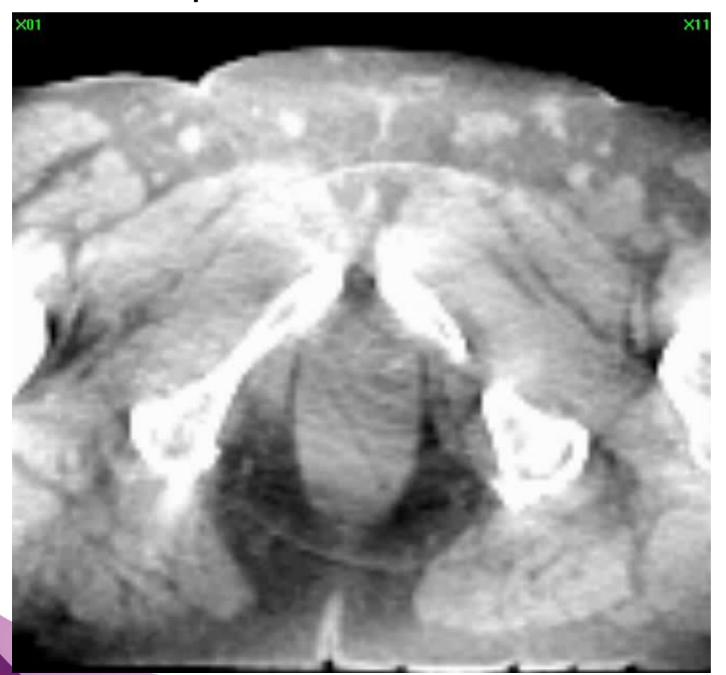
Pelvic patients using diet / drinking protocol

Immobilization and fixation:

- Head&Neck using head support
- Lung using 4D CBCT.







Reconstructed CBCT



To improve image quality:

Dietician

- Mild regimen of laxatives
- Diet

Fixed treatment times

	gas	faeces moving g	
no diet	68%	61%	45%
with diet	42%	23%	22%

- reduced percentage of faeces and gas
- reduced percentage of moving gas, hence improved image quality

Lips et al. Ijrobp 2011

- 739 patients without diet, 205 patients with diet
- Diet instructions on leaflet
- No reduction of intrafraction movement

McNair et al. 2011

- 22 patients using questionaires
- Rectal filling consistency not improved
- Diet + fixed treatment times, no laxatives

Conclusion:

- Drinking and dietery protocol are needed for clear patient communication BUT
- Won't solve the whole problem of intra/interfraction motion (additional tools are needed)



Aim of Patient preparation and positioning

Minimize the difference in patient position between simulation and treatment sessions: *inter*-fraction motion

Tools:

Patient compliance:

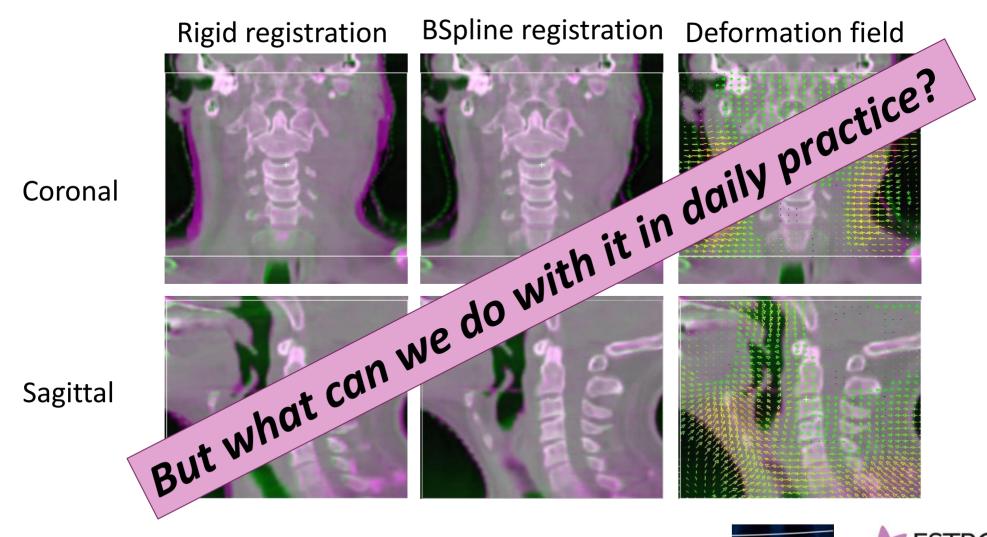
Pelvic patients using diet / drinking protocol

Immobilization and fixation:

- Head&Neck using head support
- Unfortunate differences

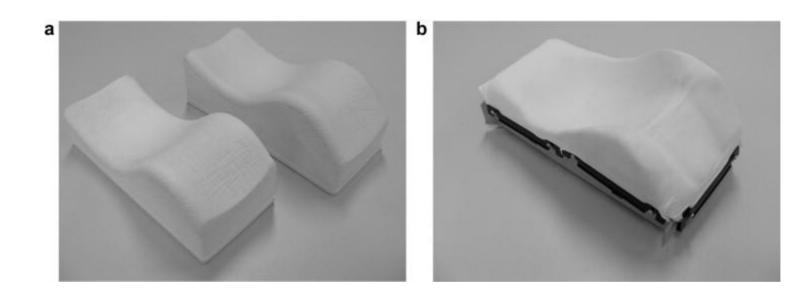


Head&Neck patients: head support





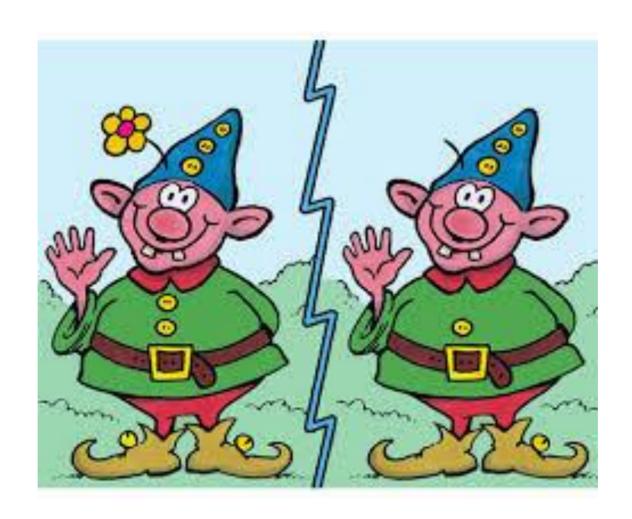
Head&Neck patients: head support



- Reduction of the average difference between fractions in set up of the bony anatomy.
- Reduction in the difference of the shape of the bony anatomy between fraction.

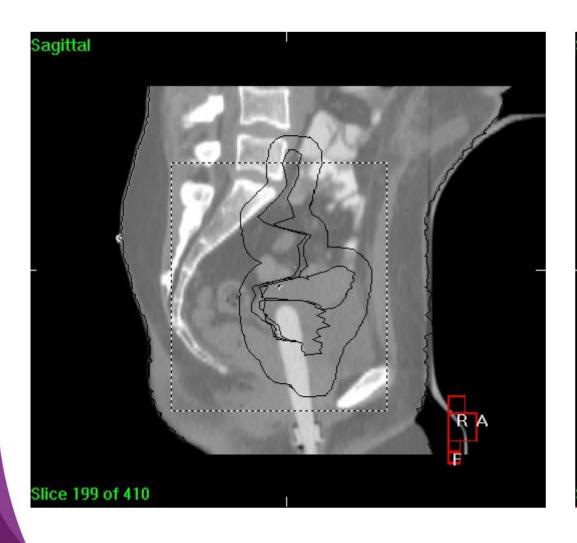
Creating unfortunate differences

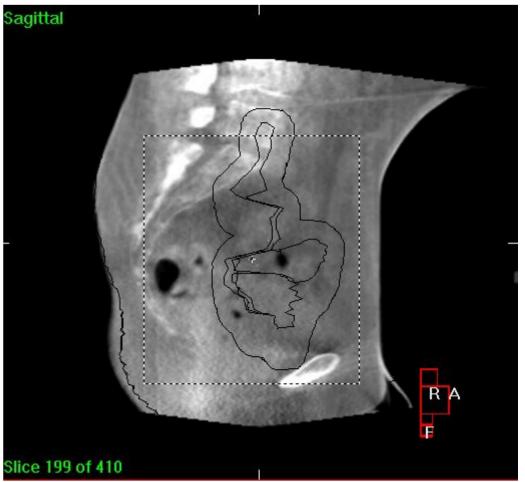
Between CT and treatment





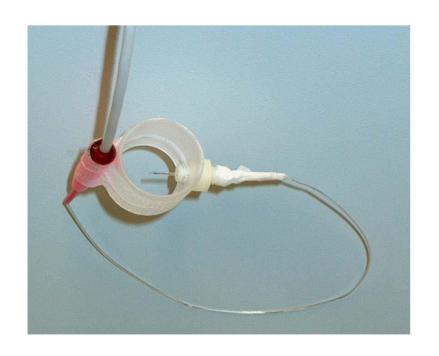
Example 1: Look for differences...

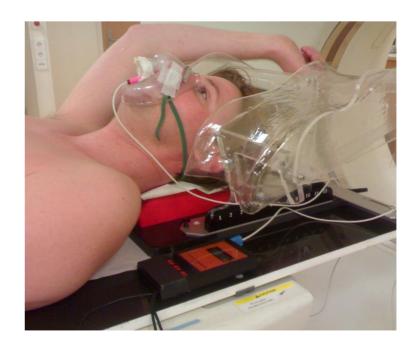






Example 2: Respiratory monitoring system





- 4D CBCT scans with and without oxygen mask
- 3D tumor motion was assessed for tumor mean position and amplitude

Respiratory monitoring system

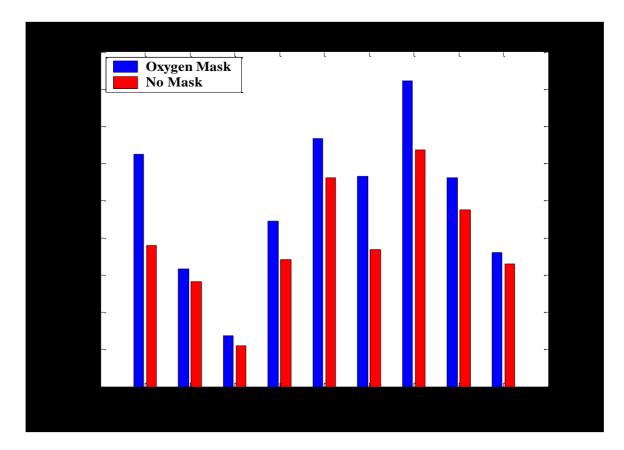
With oxygen mask

Without oxygen mask

	LR (cm)	CC	AP		LR (cm)	CC (cm)	AP (cm)
		(cm)	(cm)				
Σ	0.18	0.23	0.23	Σ	0.15	0.21	0.22
σ	0.16	0.19	0.19	σ	0.18	0.17	0.20
Mean	0.06	0.03	0.00	Mean	0.04	0.08	-0.09

No significant difference in tumour mean position

Respiratory monitoring system



M = 29%, SD = 19%, p = 0.0017

Difference in breathing amplitude!





Deformable registration decreases the need for good immobilization

A.True

B.False



Aim of Patient preparation and positioning

Minimize the difference in patient positioning during the treatment session: *intra*-fraction motion

Tools:

Increasing patient compliance:

Practical session SBRT

Immobilization and fixation:

Lung using 4D CBCT.

Practical session

In case of hypofractioned RT:

- Patient visit the linac
- Session is completely performed but no Gray's are given

Advantages:

- Patient gets acquinted with workflow
- Set-up accuracy can be assesed:
 - > is the intra# motion acceptable?
- Is it do able for the patient?
- Is the image quality sufficient?
- Precautions can be made:
 - Pain/stress relief
 - Additional margins/replanning



Hypo fractionated lung

On-line lung tumor match with CBCT: 3 x 18 Gy

(first protocol design without arc therapy and inline scanning)

Aligning the patient: 5 min

First CBCT scan: 4 min

Registration: 5 min

Manual table shift: 3 min

Second CBCT scan: 4 min

Evaluation CBCT scan: 1 min

Beam delivery: 25 min

Post treatment CBCT scan: 4 min





Antoni van Leeuwenhoek Hospital



Antoni van Leeuwenhoek Hospital

59 Patients, 3 fractions per patient

		LR (mm)	CC (mm)	AP (mm)
Residual Inter- fraction	GM	0.2	0.6	-0.6
	Σ	0.8	0.8	1.0
	σ	1.1	1.1	1.4
Intra-fraction	GM	0.0	1.0	-0.9
	Σ	1.2	1.3	1.9
	σ	1.2	1.4	1.7



Intrafraction motion is the motion of a patient within a session

- A. True
- B. False





Patient compliance won't impact intrafraction motion

A. True

B. False





Minimize the difference in patient position

- → Minimize the difference in patient position
 - 1. between simulation and treatment sessions
 - 2. during the treatment session
- Maximize the distance between target volume and organs at risk

Tools:

- Immobilization and fixation
- Patient compliance

Minimize the difference in patient position

Maximize the distance between target volume and organs at risk

Tools:

Immobilization and fixation:

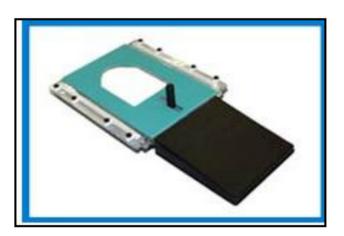
Bellyboard for pelvic patients

Patient compliance:

Breath hold for breast patients

Belly board pelvic patients



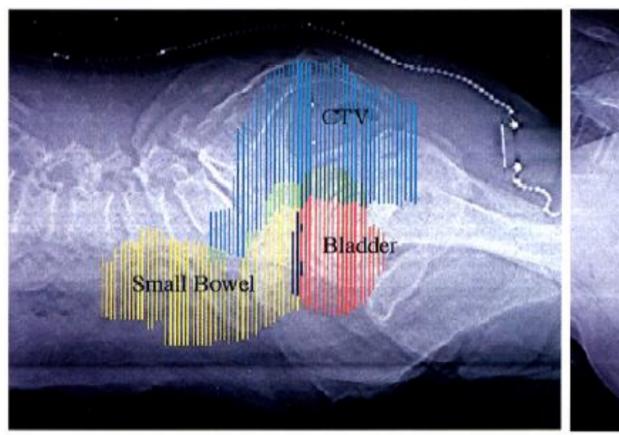


Belly board





Belly board pelvic patients



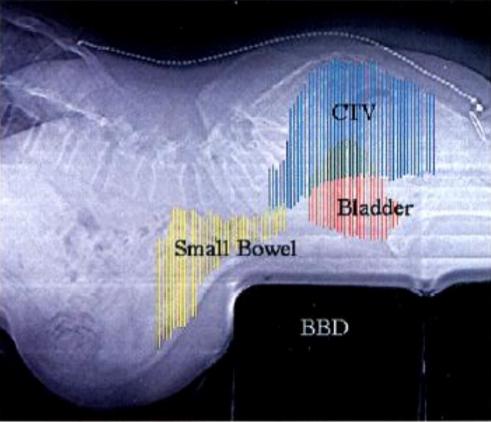


Fig. 2. Pilot localization, lateral view is shown (a) for simulation without BBD and (b) with BBD. The clinical target volume (CTV), small bowel, and bladder are shown. Note a dramatic shift in small bowel in the cephalic direction with the BBD.

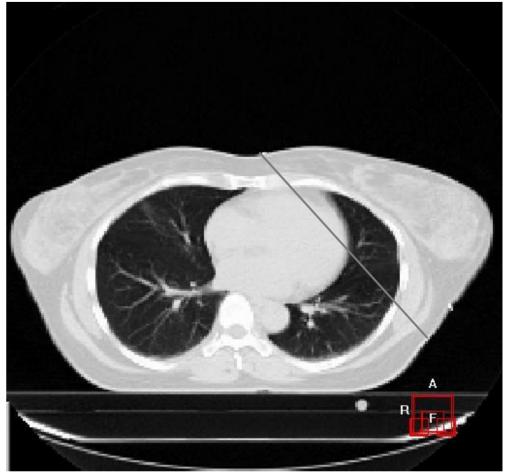


Breath hold for breast patients

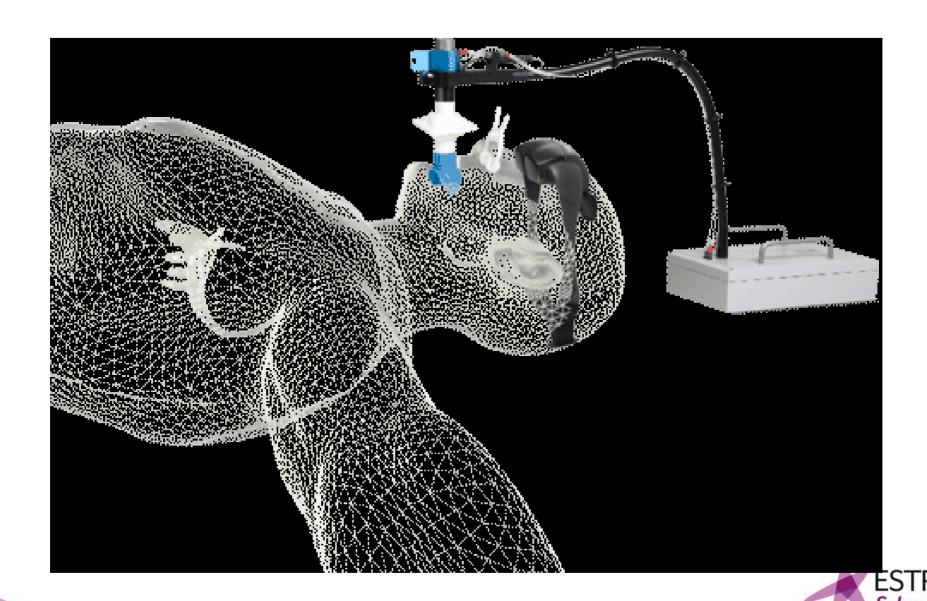
Normal inspiration

Deep inspiration



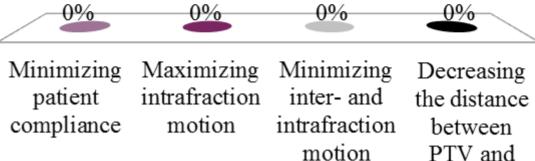


Essential: education & compliance



Patient preparation and immobilization aims at:

- A. Minimizing patient compliance
- B. Maximizing intrafraction motion
- C. Minimizing inter- and intrafraction motion
- D. Decreasing the distance between PTV and OAR's





OAR's

Conclusion

The first step in radiation therapy is to minimize

- the difference in patients anatomy and set-up between CT en treatment
- the difference in patients anatomy and set-up between treatment days

and to maximize

- patient stability
- the distance between target volume and organs at risk



Conclusion

The first step in radiation therapy is to minimize

• the difference in patients anatomy and set-up between CT en treatment

 the difference in patients anatomy and set-up between treatment days

and to maximize

- patient stability
- the distance between target volume and organs at risk

Rotations
Rotations
Deformations
Offline protocol OAR



Conclusion

https://espace.cern.ch/ULICE-results/Shared%20Documents/D.JRA 5.1 public.pdf

'Recommendations for organ depending optimized fixation systems'





Pre-treatment imaging





Mirjana Josipovic

Dept. of Oncology, Rigshospitalet & Niels Bohr Institute, University of Copenhagen Denmark

Advanced skills in modern radiotherapy
June 2017



Intended learning outcomes

- Illustrate the importance of a particular pre-treatment imaging modality for radiotherapy
- Comprehend the additional value of applying combined information from several imaging modalities for radiotherapy planning
- Identify uncertainties of pre-treatment imaging modalities



Pre-treatment imaging for radiotherapy

• CT: computed tomography

PET: positron emission tomography

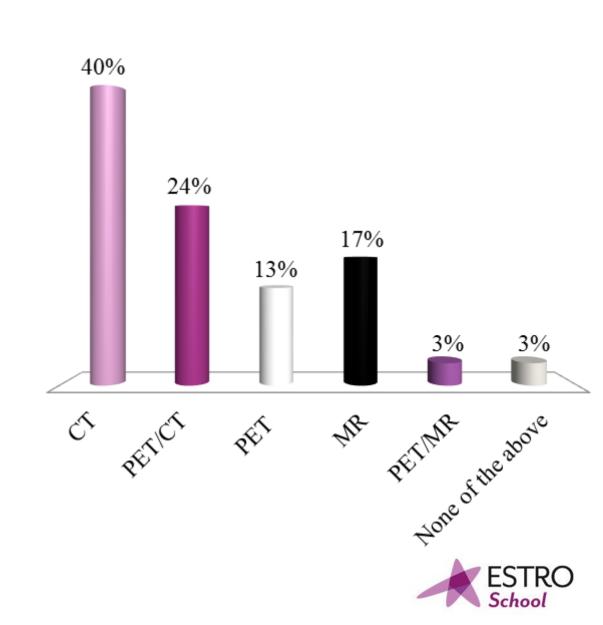
• MR: magnetic resonance



Do you have experience with...?

- A. CT
- B. PET/CT
- C. PET
- D. MR
- E. PET/MR
- F. None of the above

Multiple answers possible!



Which imaging modalities do we need for modern state of the art radiotherapy?

A. CT

B. PET

C. MR

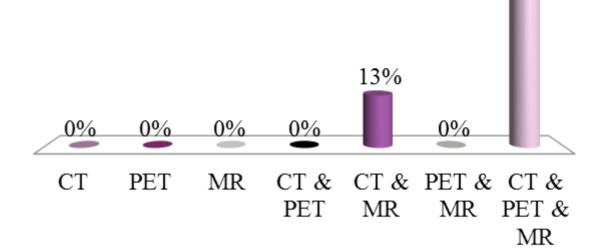
D. CT & PET

E. CT & MR

F. PET & MR

G. CT & PET & MR





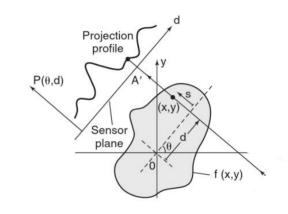


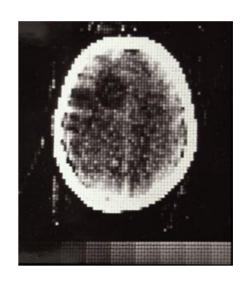
88%

CT chronology

- 1917 mathematical grounds for CT reconstruction
- 1971 first clinical CT
- 1990 spiral CT
- 1993 dual slice
- 2003 32-slice
- Today: ultrafast volume-scanning dual source, dual energy

1024x1024 matrix < 0.3 s rotation time



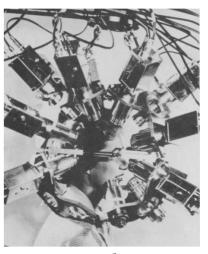


80x80 matrix 5 min rotation time

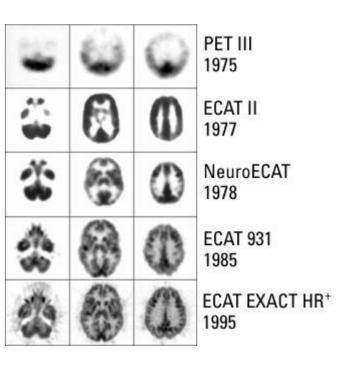


PET chronology

- 1930's radioactive tracers
- 1953/66 multidetector device
- 1975 back projection method for PET
- 1979 fluorine 18 deoxy glucose (FDG)
- 2000 PET/CT "medical invention of the year"



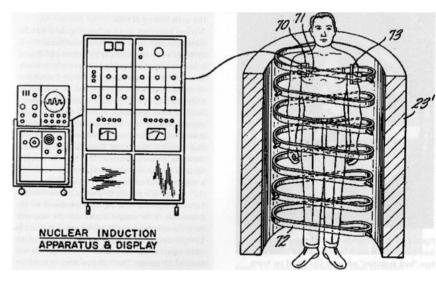
Wagner et al. 1998

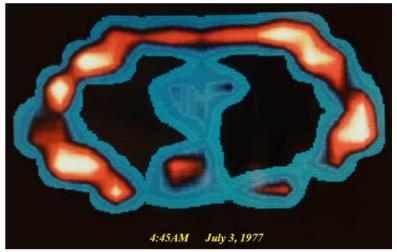




MR chronology

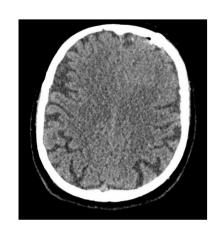
- 1937 nuclear magnetic resonance
- 1956 Tesla unit
- 1972 Damadian invention
- 1977 first MR scan
- 1993 functional MR

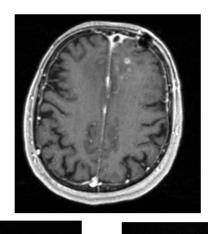


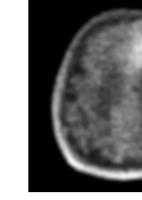


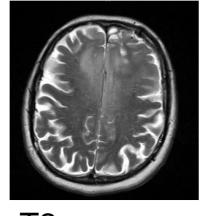


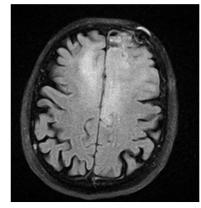
CT MR PET

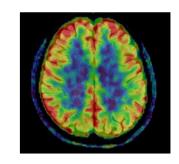






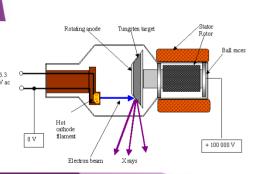


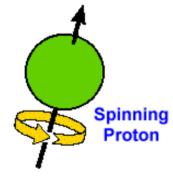


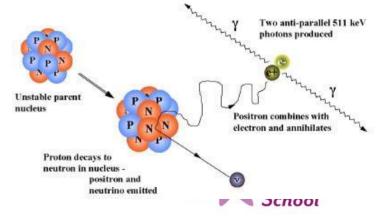


T2

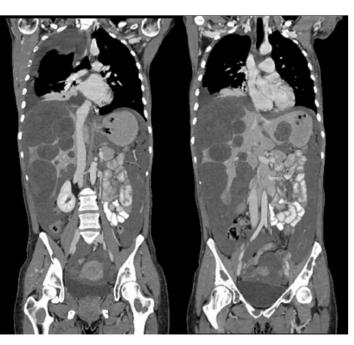
flair

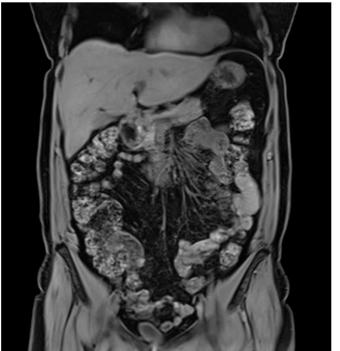


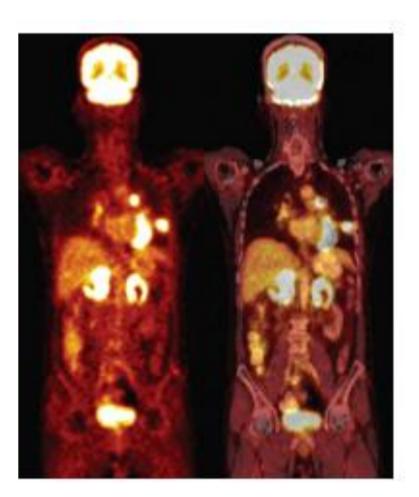




CT MR PET









What do we see?

Morphology

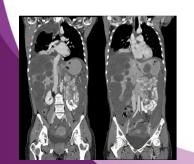
> CT, MR

(patologic) anatomy

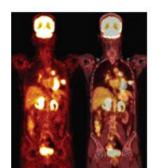
Biological processes

> PET, MR

Tumour metabolism
Perfusion
Organ function









Diagnostic imaging vs RT imaging

- Diagnostic
 - What is this?



- RT planning
 - > Where is this?





Why we need CT



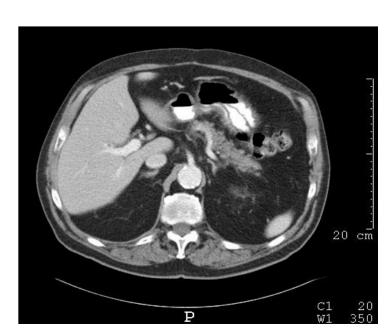
CT numbers = Hounsfield units

The grey tones on the CT image represent the attenuation in every pixel/voxel

The grey tones are expressed in Hounsfield units (HU)

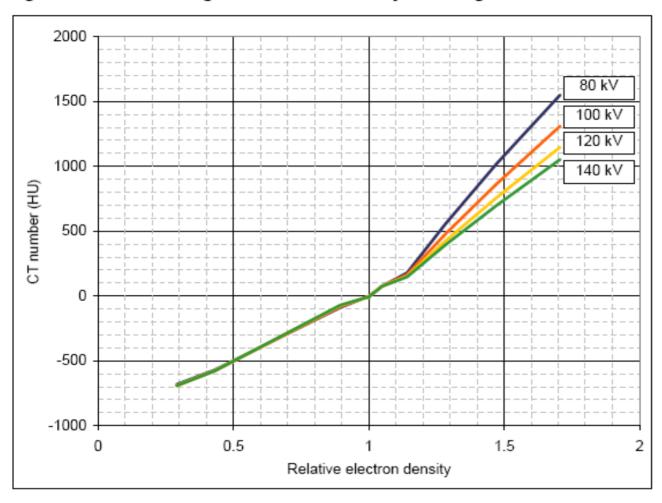
- CT numbers:

$$\begin{aligned} \mu_{obj} - \mu_{water} \\ HU = ----- x 1000 \\ \mu_{water} \end{aligned}$$



Hounsfield units → electron density

Figure 4. CT number against electron density at a range of kVs



Necessary for dose calculation

Calibration curve needed for each applied kV



How well can we trust the imaging information?





Image artifacts

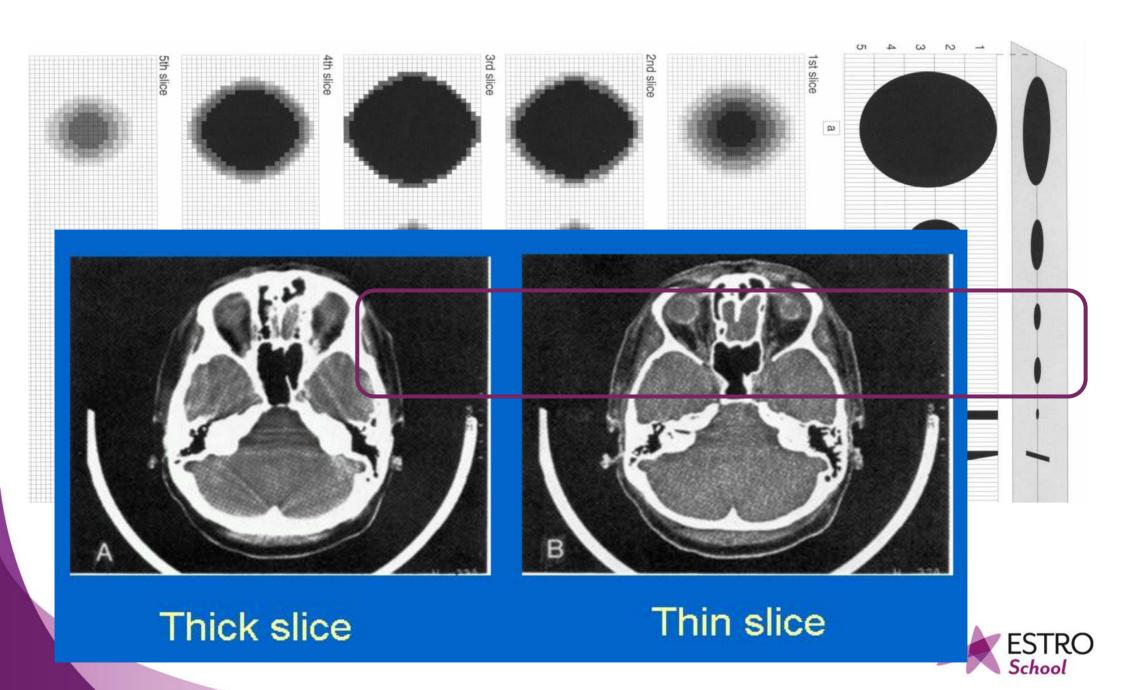
Definition:

Systematic deviation between the HU in the reconstructed image and the objects correct attenuation's coefficient

- Partial volume artefacts
- Streak artefacts
- Ring artefacts
- Motion artefacts
- Noise



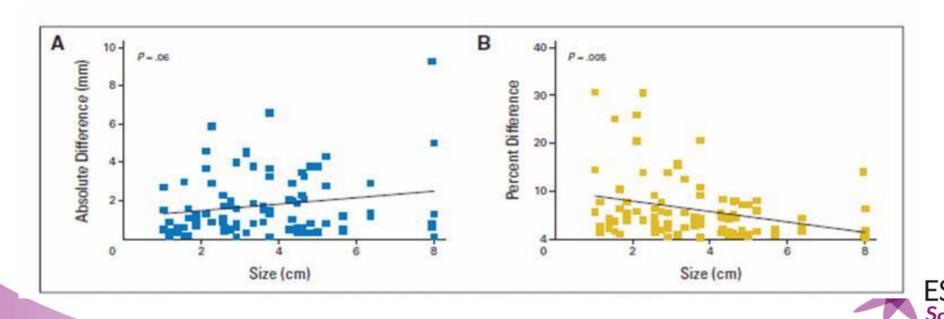
Partial Volume artefacts



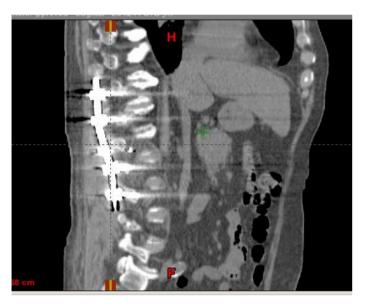
Variability of Lung Tumor Measurements on Repeat Computed Tomography Scans Taken Within 15 Minutes

Size of Tumor (cm)	Standard Deviation (mm)	Example Turnor		
		Size (cm)	Range As a Result of Variability (cm)*	% Change As a Result of Variability
1-3	2.0	2	1.6-2.4	± 20
3-5	2.3	4	3.5-4.5	± 12
5-8	3.3	7	6.3-7.7	± 9

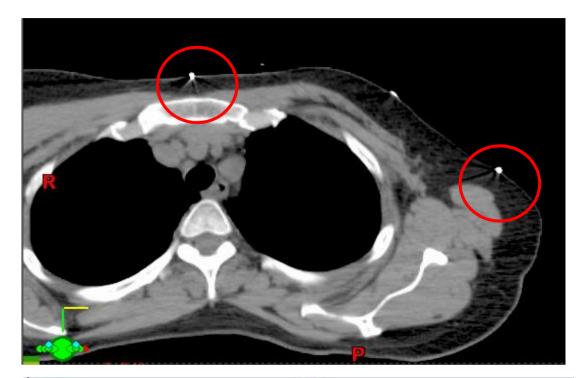
For a lesion measuring 4 cm, CT variability can lead to measurements from 3.5 to 4.5 cm

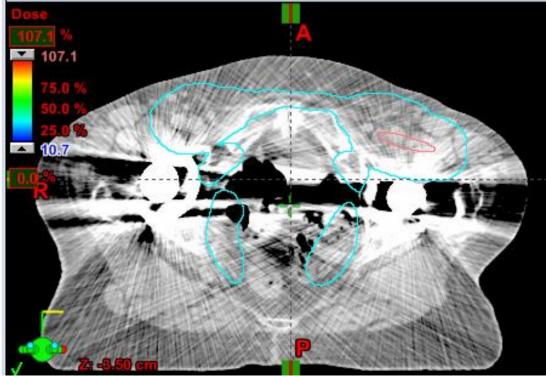


Streak artefacts









Metal artifact reducton sw

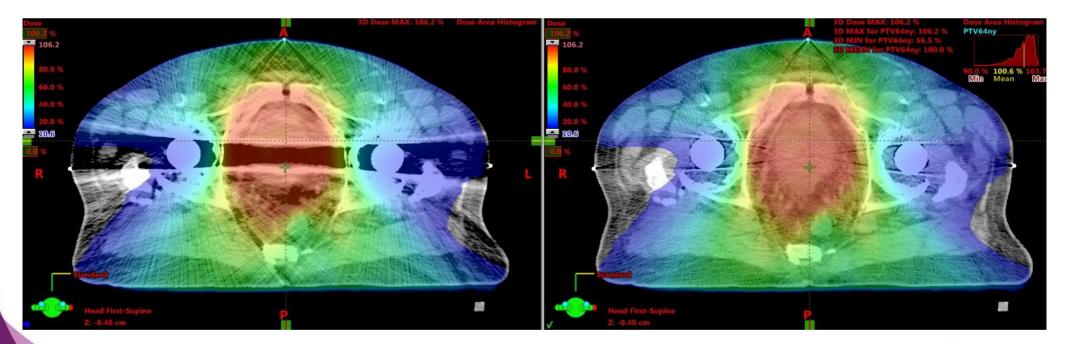
- Dual Energy CT (DECT)
 - Used two different X-ray energies
 - "Virtual monochromatic" scans
- Iterative metal artifact reduction software
 - MAR, iMAR, O-MAR...



MAR - impact on dose planning

Dose calculation for 10 patients with iMAR

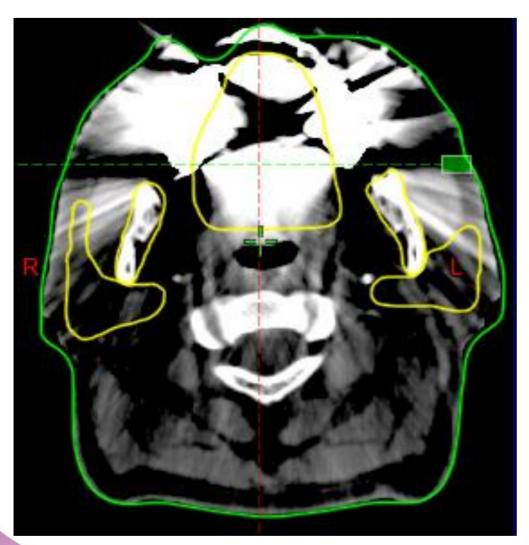
No difference in dose compared to manual override

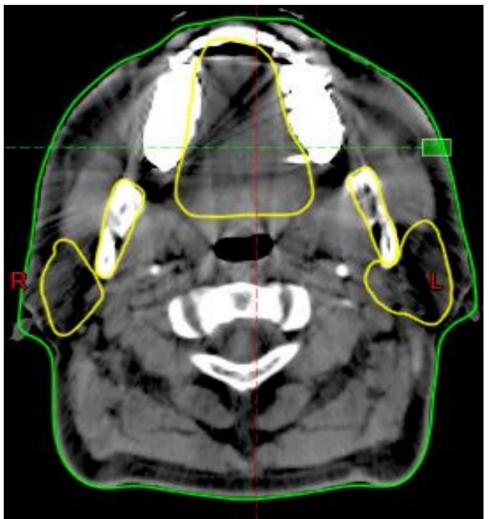




MAR- impact on contouring

Head and neck contouring by a radiation oncologist

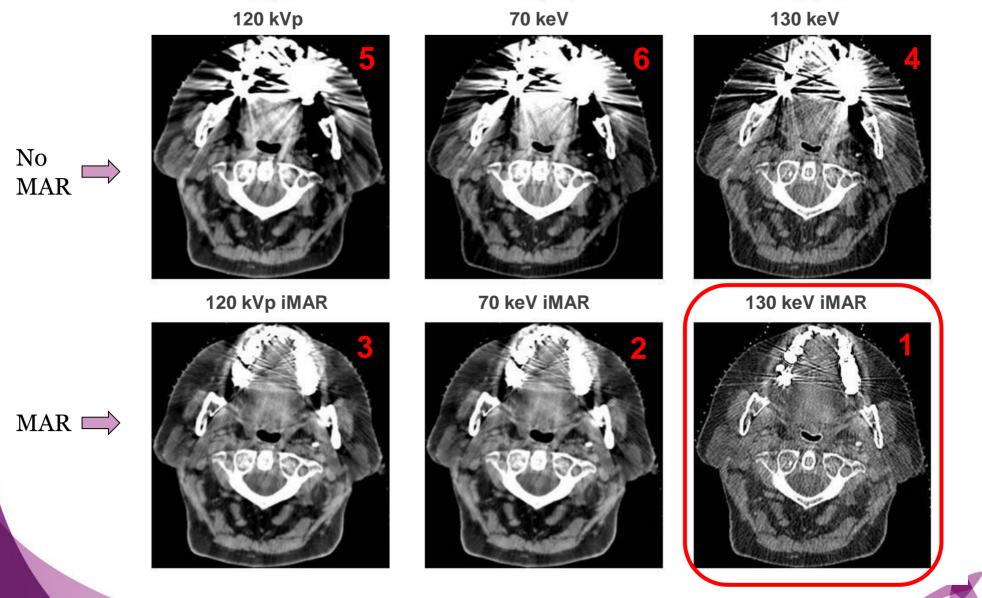






MAR combined with dual energy scan

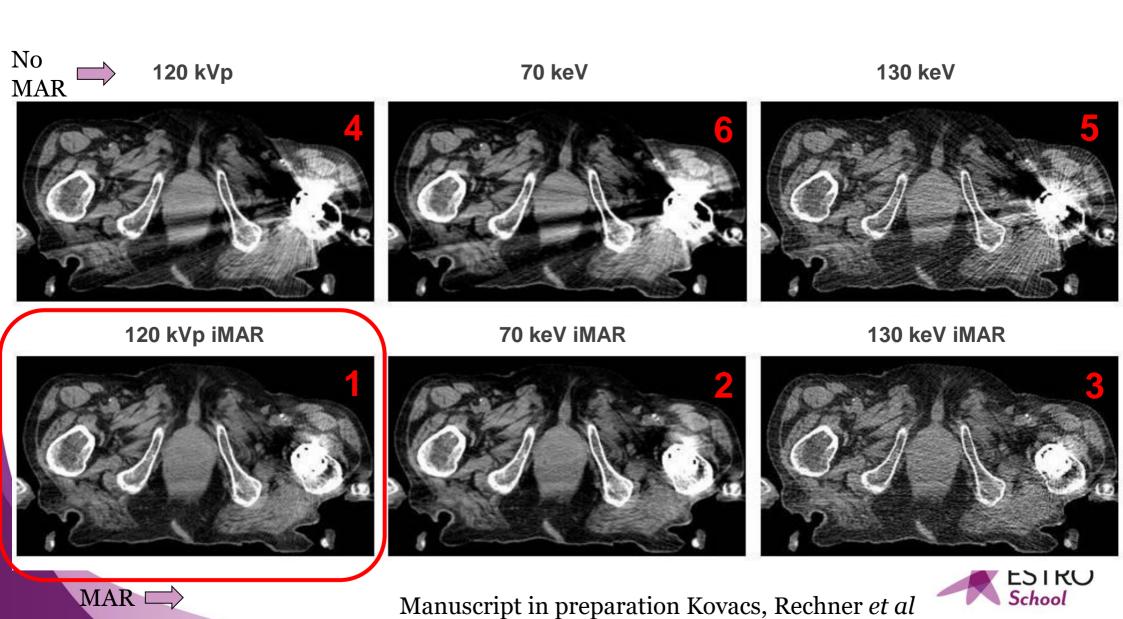
Which images do radiologists & oncologists prefer?



Manuscript in preparation Kovacs, Rechner et al

MAR combined with dual energy scan

Which images do radiologists & oncologists prefer?



Imaging for RT planning

- Has to be precise
- Has to provide safe judgment of the extent of the disease
- CT images are base for treatment planning

BUT

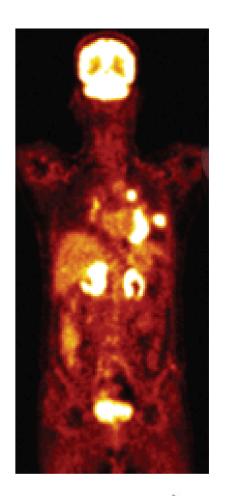
- On CT, it can be difficult to discriminate vital tumour tissue from scar tissue, oedema, atelectasis, surrounding soft tissu...
- CT can not stage correctly
 - detect small metastases
 - detect distant metastases



Added value of PET CT for radiotherapy

- Improved delineation consistency
- Improved staging



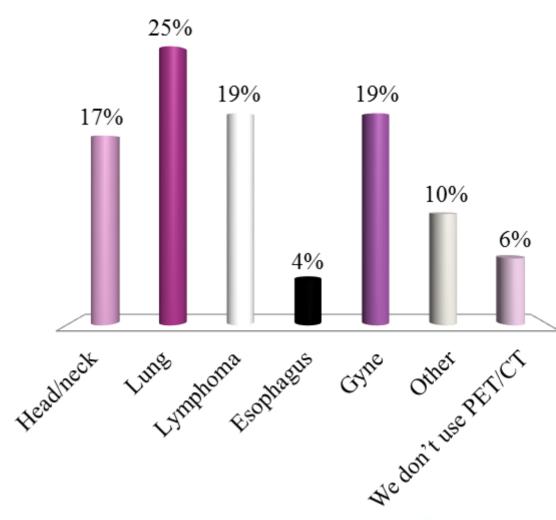




Which sites do you plan with PET/CT?

- A. Head/neck
- B. Lung
- C. Lymphoma
- D. Esophagus
- E. Gyne
- F. Other
- G. We don't use PET/CT

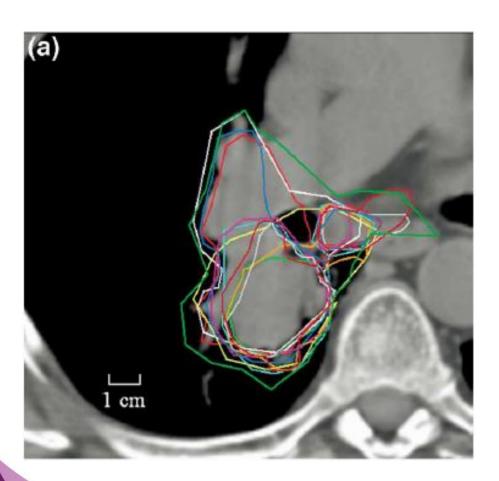
Multiple answers possible!



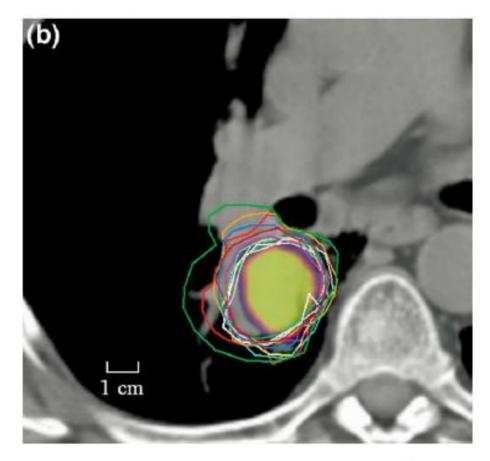


Improved delineation consistency

CT based



PET/ CT based

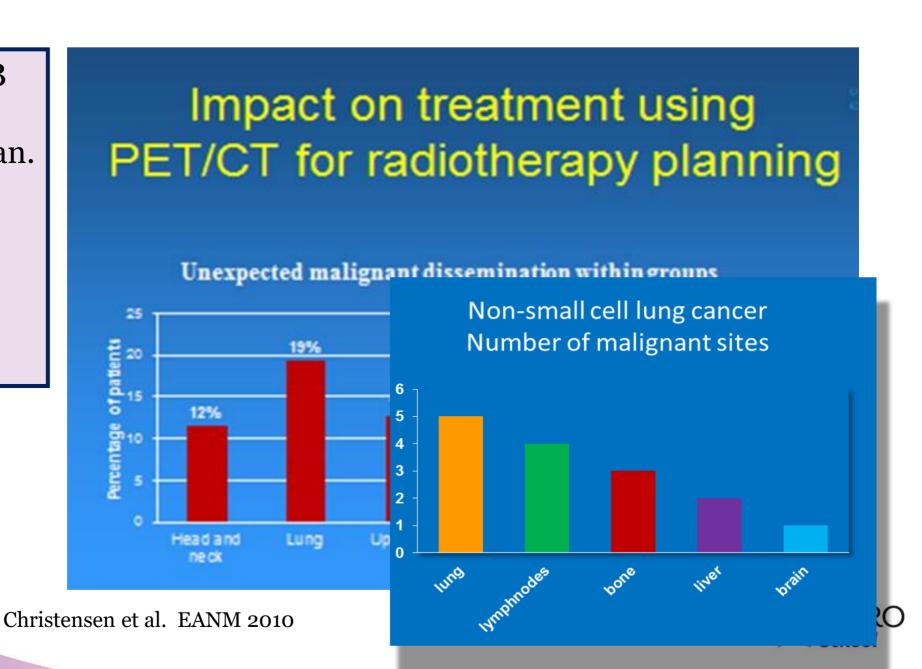




Improved staging

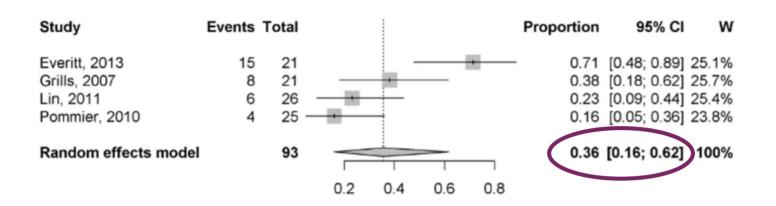
Always WB PET/CT at therapy scan.

Changing treatment strategy!

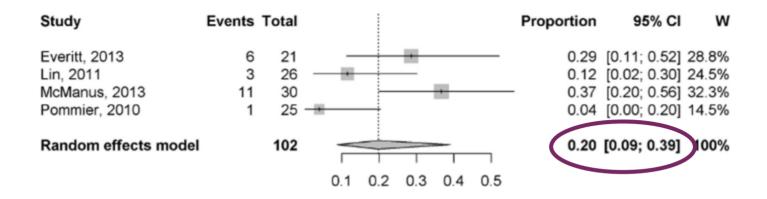


Impact of PET in lung cancer RT

Staging PET not available



Change in target definition

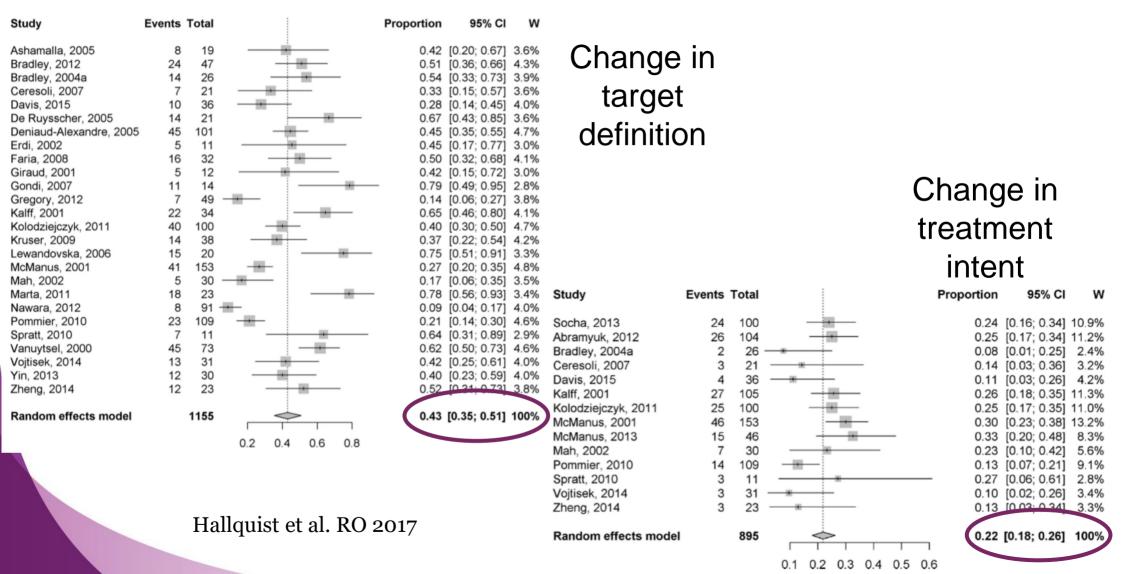


Change in treatment intent



Impact of PET in lung cancer RT

Staging PET available



Impact of PET in lung cancer RT

- Change in target definition: in 2 out of 5 patients
- Change in treatment intent: in 1 out of 5 patients

Radiotherapy and Oncology 123 (2017) 71-77



Contents lists available at ScienceDirect

Radiotherapy and Oncology





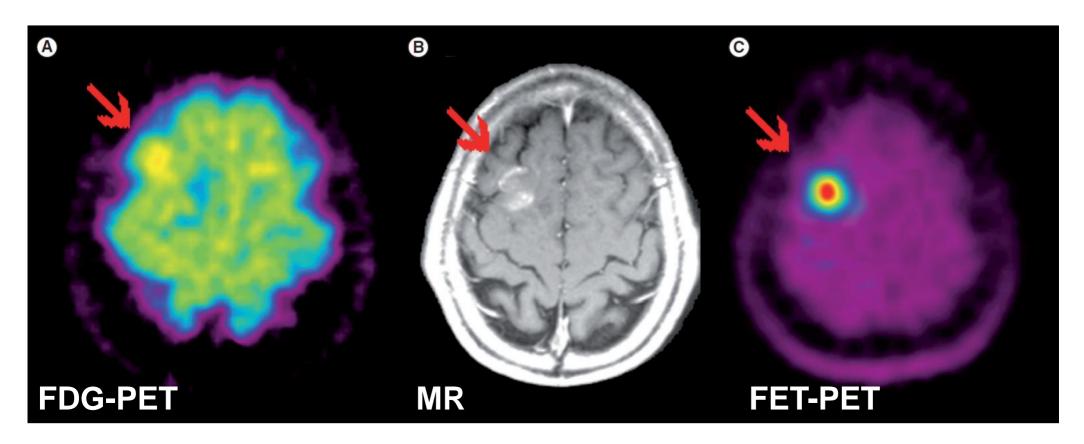
Systematic review

Positron emission tomography and computed tomographic imaging (PET/CT) for dose planning purposes of thoracic radiation with curative intent in lung cancer patients: A systematic review and meta-analysis





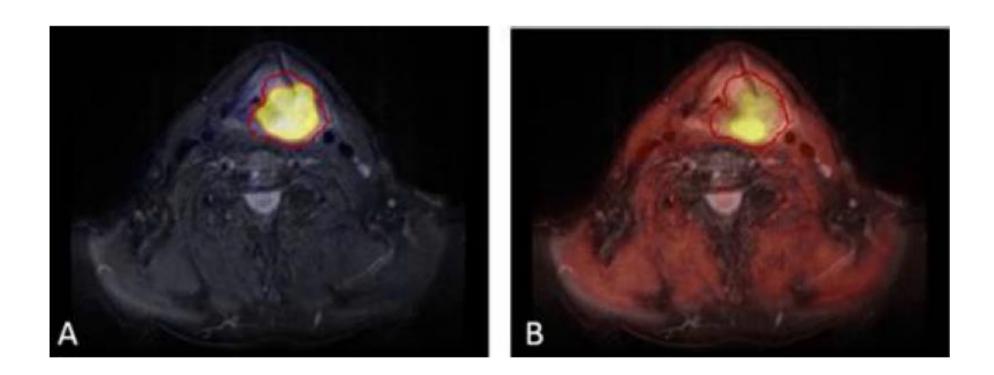
PET imaging of brain tumours



• 18F-Fluoro-Ethyl-Tyrosin (FET), aminoacid uptake



PET imaging of hypoxia with FMISO

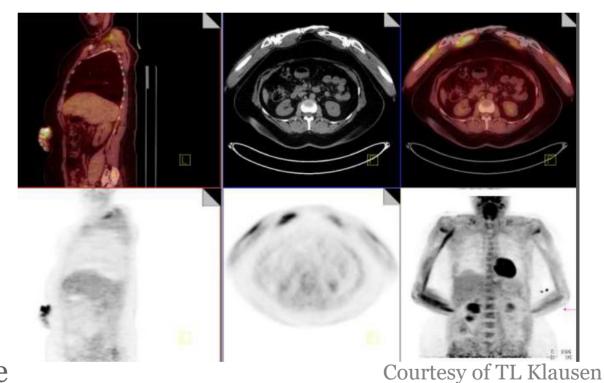


• Hypoxia area is associated with high risk of locoregional failure



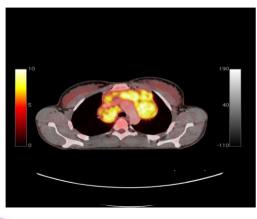
Pitfalls

- FDG is not specific
 - Not all "hot-spots" are malignant

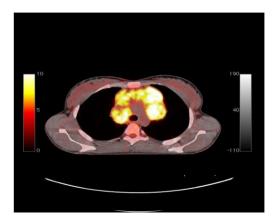


- Motion blurs the FDG uptake
 - Is it a small lesion, with high degree of motion and high SUV uptake?
 - > Is it a large lesion, without motion and low SUV uptake?

Free breathing



Breath hold



Courtesy of M Aznar



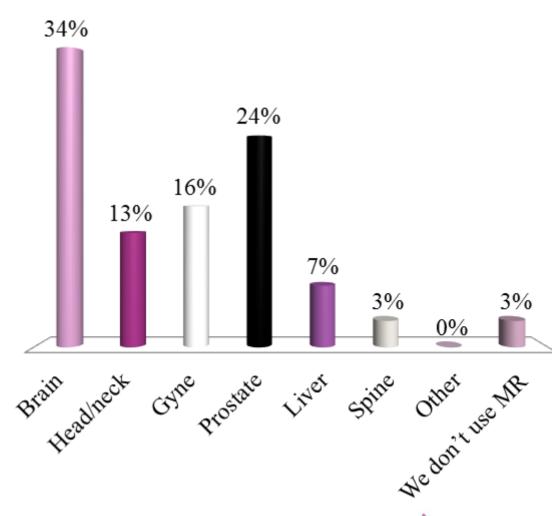
Added value of MR imaging for RT

• Superior soft tissue contrast



Which sites do you plan with MR?

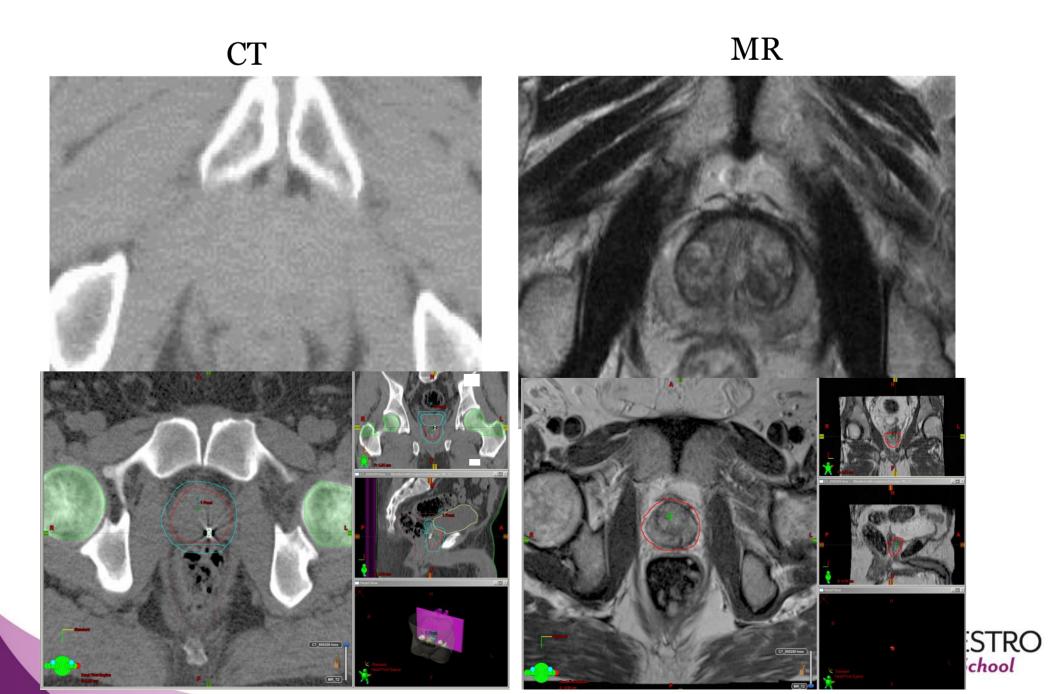
- A. Brain
- B. Head/neck
- C. Gyne
- D. Prostate
- E. Liver
- F. Spine
- G. Other
- H. We don't use MR



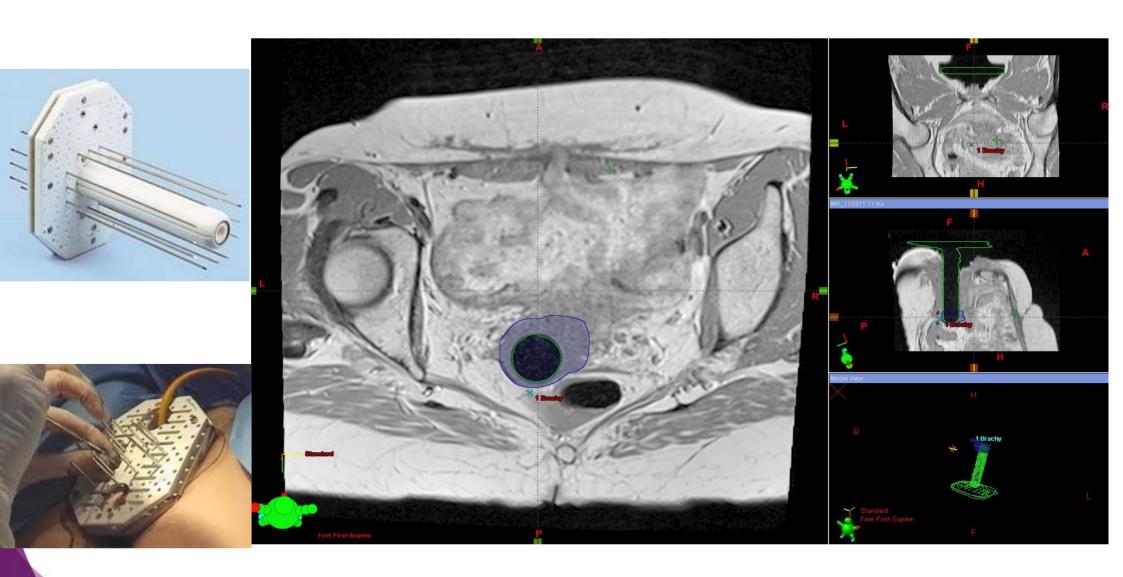
Multiple answers possible!



Prostate cancer



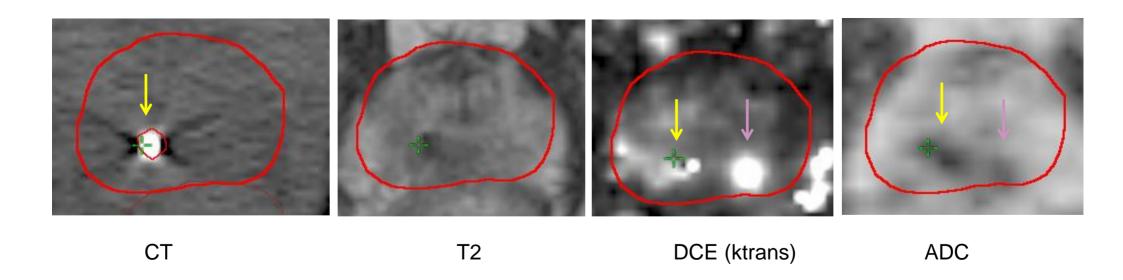
Cervix cancer - brachytherapy



dummy template for interstitial brachytherapy



Functional imaging with MR



DCE = dynamic contrast enhanced

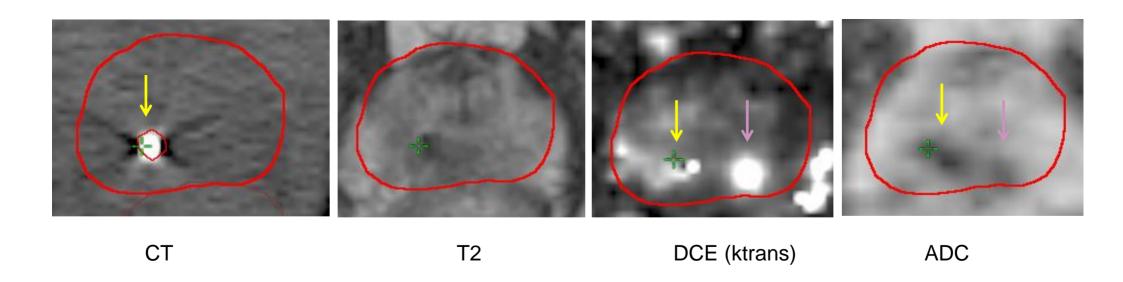
• high signal due to increase in capilar permeability

ADC = apparent diffusion coefficient

lack of signal due to high cell density



Functional imaging with MR



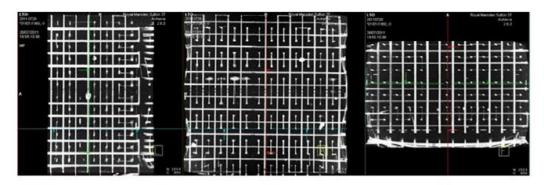
Potential biomarker for prostate cancer progression

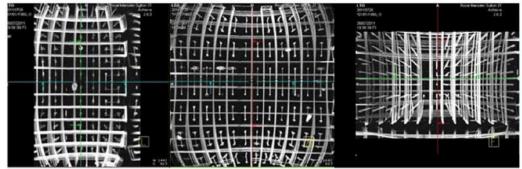
- dose escalation
- no compromises in treatment plan



Pitfalls

Geometric distortion





Schmidt & Payne PMB 2015

- No direct relation with electron density
 - > CT atlas corregistration
 - > MR segmentation



PET/MR for radiotherapy?



European Journal of Kadiology 84 (2015) 1285-1292



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journal homepage: www.elsevier.com/locate/ejrad



Oncological whole-body staging in integrated ¹⁸F-FDG PET/MR: Value of different MR sequences for simultaneous PET and MR reading



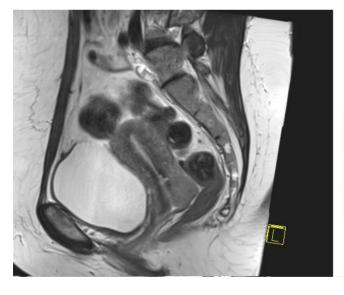
Benedikt M. Schaarschmidt^{a,b}, Johannes Grueneisen^b, Philipp Heusch^{a,*}, Benedikt Gomez^c, Karsten Beiderwellen^b, Verena Ruhlmann^c, Lale Umutlu^b, Harald H. Quick^{d,e}, Gerald Antoch^a, Christian Buchbender^a

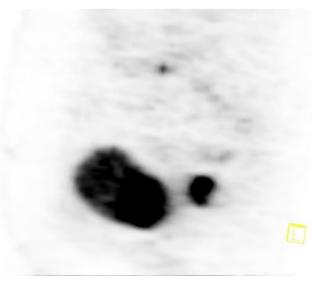
Conclusion: In conclusion, T2, TIRM, and contrast-enhanced T1 provide a high quality of lesion detectability and anatomical allocation of FDG-avid foci. Their performance is at least comparable to contrast-enhanced PET/CT. Non-enhanced T1 may be omitted and the necessity of DWI should be further investigated for specific questions, such as assessment of the liver.

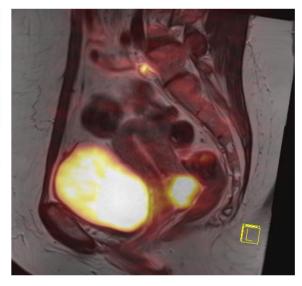


PET/MR

Images courtesy of AK Berthelsen







T2 sag (MR)

FDG-PET

PET/MR

Table 3 Difference in millimeters between the registrations of lesion isocenters measured in three axis with PET/MR

Axis	T2-weighted, PET [mean (SD)] (mm) (25 lesions)	ADC, PET [mean (SD)] (mm) (20 lesions)
x	2.28 (1.36)	4.15 (2.27)
y	2.58 (1.97)	2.08 (1.52)
Z	2.95 (2.25)	2.15 (1.78)
Total difference	5.22 (1.97)	5.79 (1.70)

ADC, apparent diffusion coefficient.

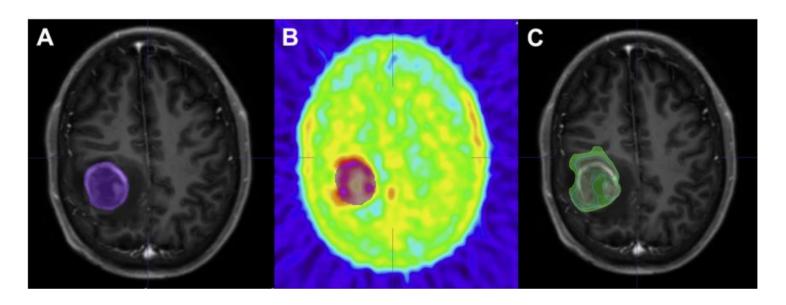
Table 6 Overlap between T2-weighted images and PET, and between ADC and PET

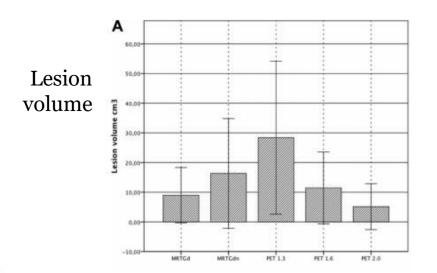
Tumors	T2-weighted, PET [mean (SD)]	ADC, PET [mean (SD)]
All (ml)	0.64 (0.13)	0.56 (0.14)
< 14 ml	0.51 (0.13)	0.44 (0.14)
14-62 ml	0.63 (0.10)	0.58 (0.11)
≥62 ml	0.76 (0.03)	0.66 (0.04)

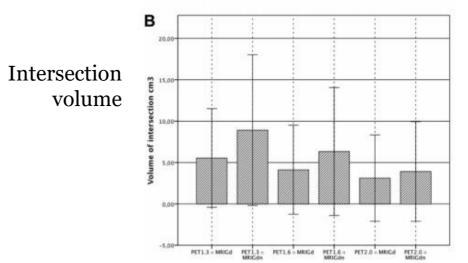
ADC, apparent diffusion coefficient.



PET/MR imaging of brain tumours





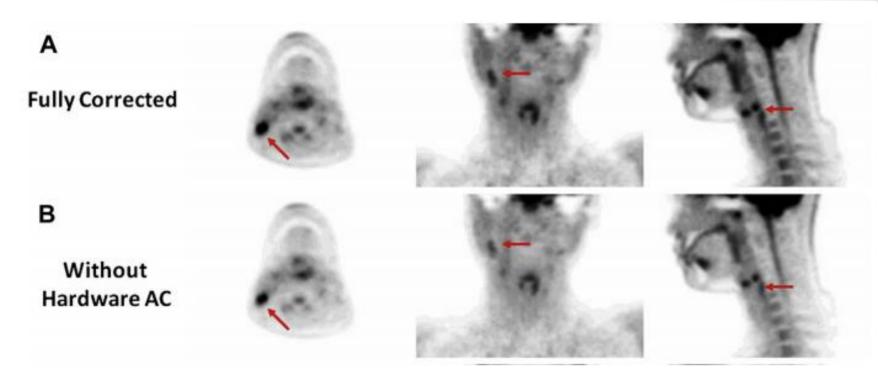




PET/MR pitfall

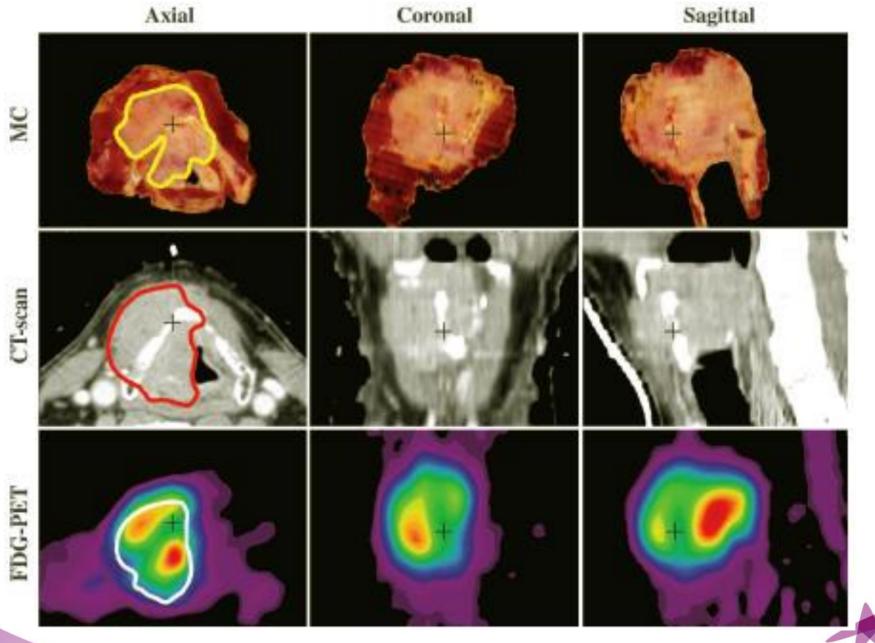
100 FURO

• MR coils impair PET signal





Challenge of multi modality imaging



Conclusion (1)

- Illustrate the importance of a particular pre-treatment imaging modality for radiotherapy
 - > CT is needed for calculation of dose distribution
 - PET adds value for staging, distinguishing tracer avid areas/volumes
 - > MR increased soft tissue contrast



Conclusion (2)

- Comprehend the additional value of applying combined information from several imaging modalities for radiotherapy planning
 - More reproducible target definition
 - More precise target definition
 - Optimal treatment strategy



Conclusion (3)

- Identify uncertainties of pre-treatment imaging modalities
 - Artefacts in images
 - Differences in (spatial) info on each modality





TARGET VOLUME DELINEATION



Sofia Rivera, MD, PhD Radiation Oncology Department Gustave Roussy Villejuif, France

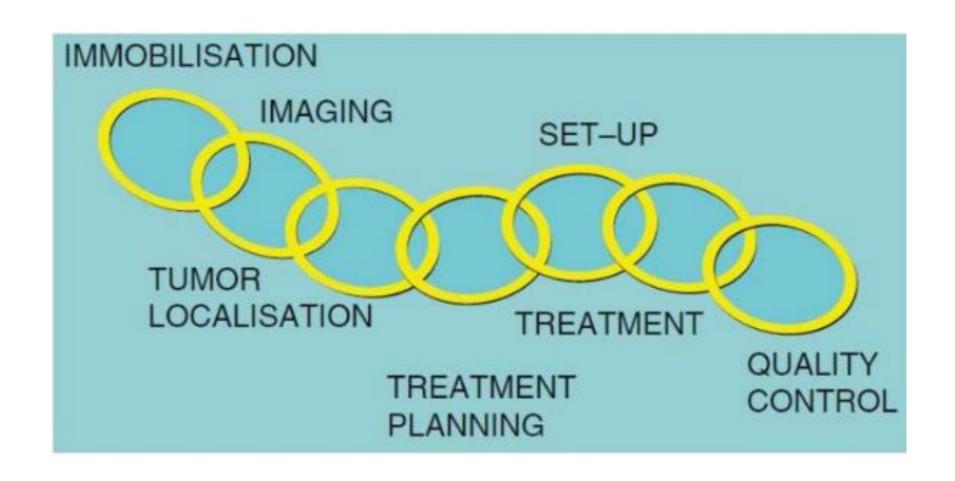


Advanced skills in modern radiotherapy June 11, 2017

Learning outcomes

- Understand why heterogeneity in contouring is a major weak point in modern radiotherapy
- Discuss the challenges in contouring target volumes
- Identify skills required to delineate target volumes
- Identify tools for improving learning in delineation
- Identify adequate imaging modalities according to the target to delineate
- Discuss the impact and consequences of inaccurate delineation of target volumes

Delineation: one of the links in the treatment chain





Why is delineation important?

- Radiotherapy planning is nowadays mostly based on CT scans
- Constraints for dose distribution are used
- DVH are calculated based on the contours
- Field arrangements are becoming more complex
- An error in contouring will therefore translate in a systematic error all along the treatment and may have consequences:
 - Jeopardizing treatment efficacy
 - Impacting treatment toxicity



Do we need to improve?



Published in final edited form as:

Radiother Oncol. 2012 April: 103(1): 92-98. doi:10.1016/j.radonc.2012.02.010.

Heterogeneity in head and neck IMRT target design and clinical practice

Theodore S. Hong^a, Wolfgang A. Tomé^{b,c,d}, and Paul M. Harari^{b,*}

Abstract

Purpose—To assess patterns of H&N IMRT practice with particular emphasis on elective target delineation.

Materials and methods—Twenty institutions with established H&N IMRT expertise were solicited to design clinical target volumes for the identical H&N cancer case. To limit contouring variability, a primary tonsil GTV and ipsilateral level II node were pre-contoured. Participants were asked to accept this GTV, and contour their recommended CTV and PTV. Dose prescriptions, contouring time, and recommendations regarding chemotherapy were solicited.

Results—All 20 institutions responded. Remarkable heterogeneity in H&N IMRT design and practice was identified. Seventeen of 20 centers recommended treatment of bilateral necks whereas 3/20 recommended treatment of the ipsilateral neck only. The average CTV volume was 250 cm³ (range 37–676 cm³). Although there was high concordance in coverage of ipsilateral neck levels II and III, substantial variation was identified for levels I, V, and the contralateral neck. Average CTV expansion was 4.1 mm (range 0–15 mm). Eight of 20 centers recommended chemotherapy (cisplatin), whereas 12/20 recommended radiation alone. Responders prescribed on average 69 and 68 Gy to the tumor and metastatic node GTV, respectively. Average H&N target volume contouring time was 102.5 min (range 60–210 min).

Conclusion—This study identifies substantial heterogeneity in H&N IMRT target definition, prescription, neck treatment, and use of chemotherapy among practitioners with established H&N IMRT expertise. These data suggest that continued efforts to standardize and simplify the H&N IMRT process are desirable for the safe and effective global advancement of H&N IMRT practice.



How can we answer that need?

- Adequate imaging, training and use of contouring guidelines are the main strategies to minimize delineation uncertainties (Petrič et al 2013)
- Establishing and using consensus and guidelines have shown to reduce heterogeneity in contouring

Table III. Mean and ranges of DSC before and after consensus.

Volume	Consensus volume (ml)	Mean DSC (range) Before consensus	Mean DSC (range) After consensus	
Breast	1247	0.93 (0.89-0.96)	0.95 (0.93-0.96)	
Boost	40	NA	0.75 (0.60-0.89)	
Internal mammary LN	15	0.59 (0.32-0.72)	0.71 (0.63-0.81)	
Axillary LN level I	108	0.65 (0.59-0.75)	0.70 (0.60-0.77)	
Axillary LN level II	32	0.56 (0.35-0.69)	0.76 (0.67-0.84)	
Axillary LN level III	17	0.56 (0.39-0.73)	0.74 (0.66-0.82)	
Periclavicular LN	47	0.41 (0.34-0.56)	0.56 (0.43-0.73)	
Interpectoral LN	33	0.54 (NA)	0.66 (0.55-0.78)	
Heart	731	0.91 (0.88-0.94)	0.94 (0.90-0.96)	

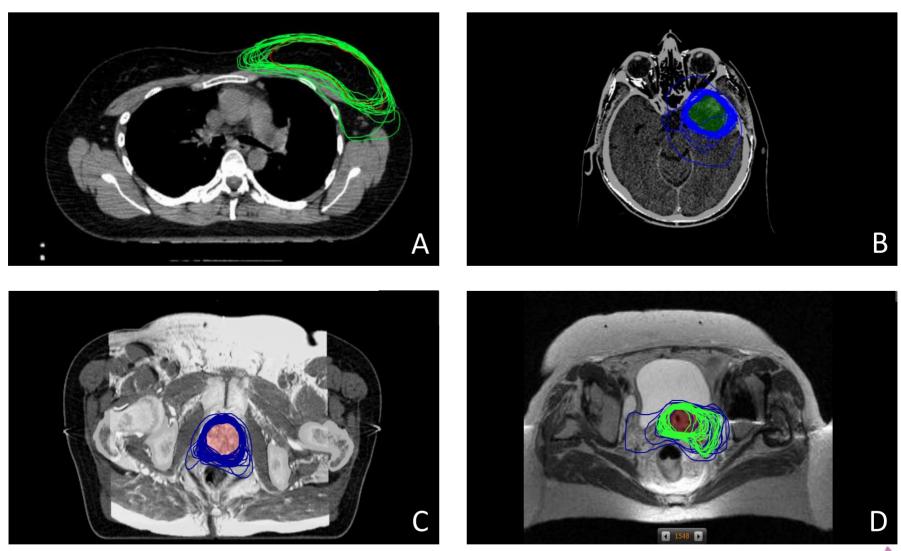
DSC, Dice similarity coefficient; NA, not available.



Inter-observer variability in contouring

Examples of participant contours from ESTRO FALCON workshops.

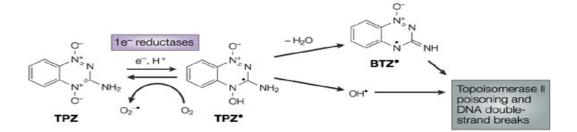
A: CTV breast, B: GTV Brain tumour, C: CTV prostate and D: GTV cervix cancer





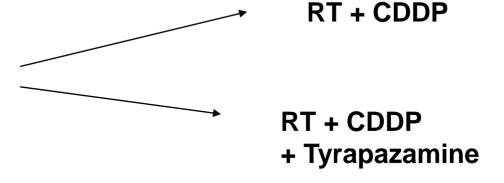
Does heterogeneity in RT matters?

- Bioreductive agent
- Radiosensitizer in hypoxia



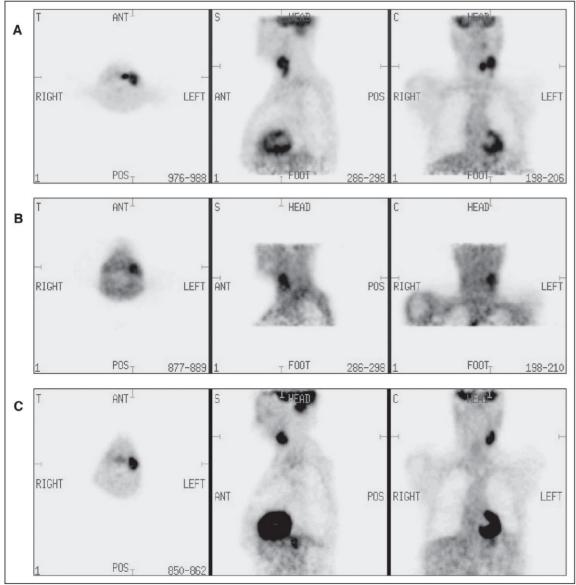
Nature Reviews | Cancer

Multicentric international Randomized phase III 853 locally advanced H&N patients





Hypoxia radioresistance



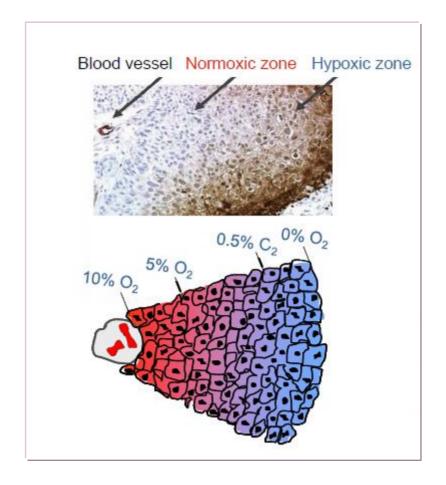
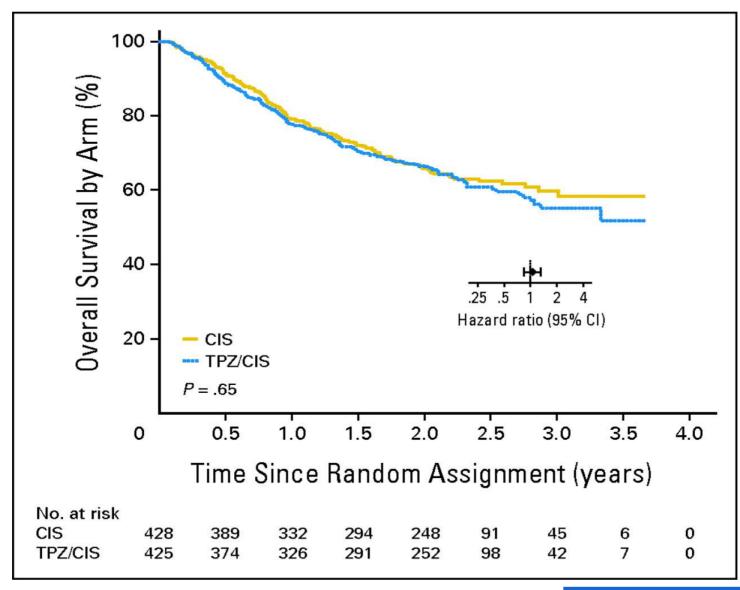




Fig 1. (A) Baseline [18F]-fluorodeoxyglucose (FDG) positron emission tomography (PET) of patient with T2N2b squamous cell carcinoma of the pyriform fossa with left nodal mass. (B) [18F]-fluoromisonidazole (FMISO) -PET at baseline, nonhypoxic primary tumor, and hypoxic node. (C) FDG-PET 12 weeks after chemoboost, complete response in nonhypoxic primary tumor, and poor response in hypoxic node. Residual tumor in nodal mass was confirmed pathologically after neck dissection.

No benefit in overall survival

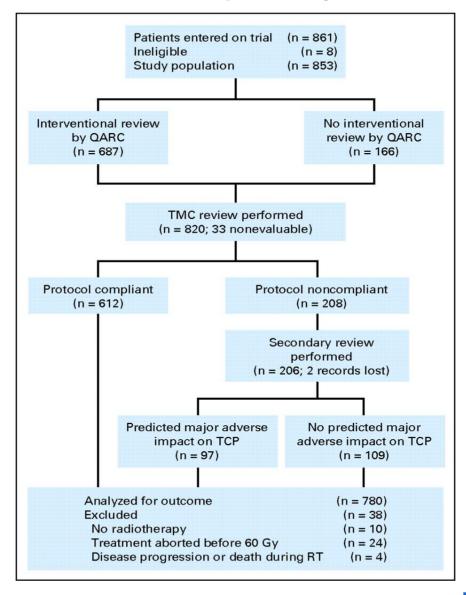


Rischin D et al. JCO 2010;28:2989-2995

JOURNAL OF CLINICAL ONCOLOGY



But... Trial quality control

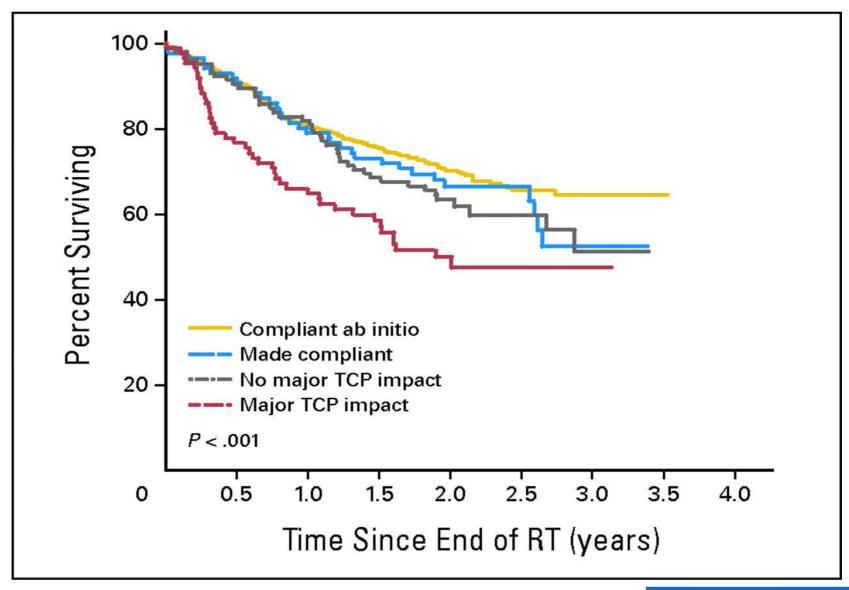


Peters L J et al. JCO 2010;28:2996-3001

JOURNAL OF CLINICAL ONCOLOGY



Impact of radiotherapy quality



Peters L J et al. JCO 2010;28:2996-3001

JOURNAL OF CLINICAL ONCOLOGY



How to improve?

- Need for a common language: ICRU
- Need for delineation guidelines and anatomical knowledge
- No absolute truth so need to specify according to which guidelines we contour
- Heterogeneity in understanding/interpreting the guidelines
- Need for teaching in contouring
- Need for evaluation in contouring



ICRU Guidelines (ICRU50): volume definition

- Volumes defined prior/ during treatment planning:
 - Gross Tumor Volume (GTV)
 - Clinical Target Volume (CTV)
 - Planning Target Volume (PTV)
 - Organs At Risk (OAR)
 - > Treated Volume
 - Irradiated Volume
- Volumes might be redefined during treatment for adaptive RT

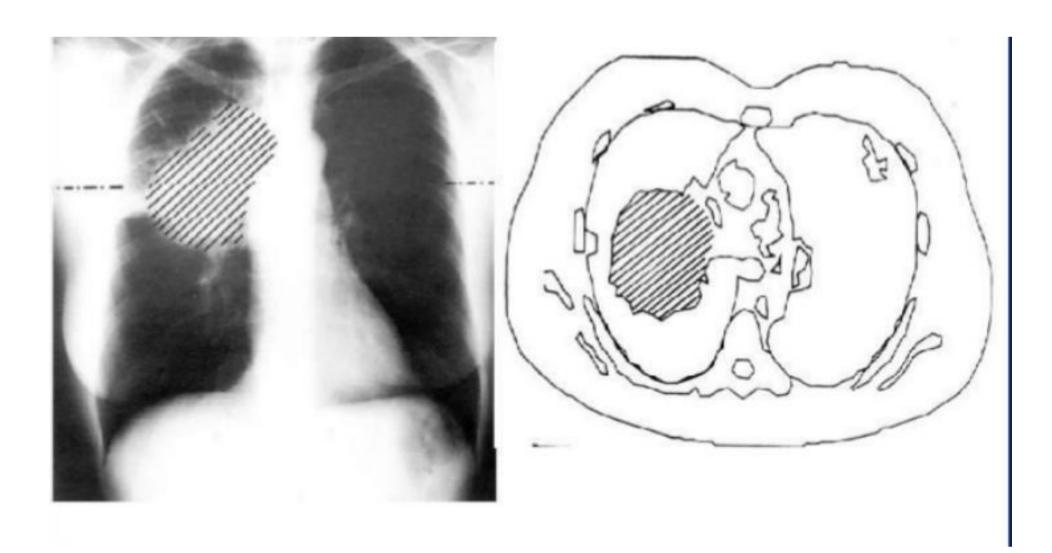


- Macroscopic tumor volume visible or palpable
- Includes:
 - Primary tumor
 - Macroscopically involved lymph nodes
 - Metastases



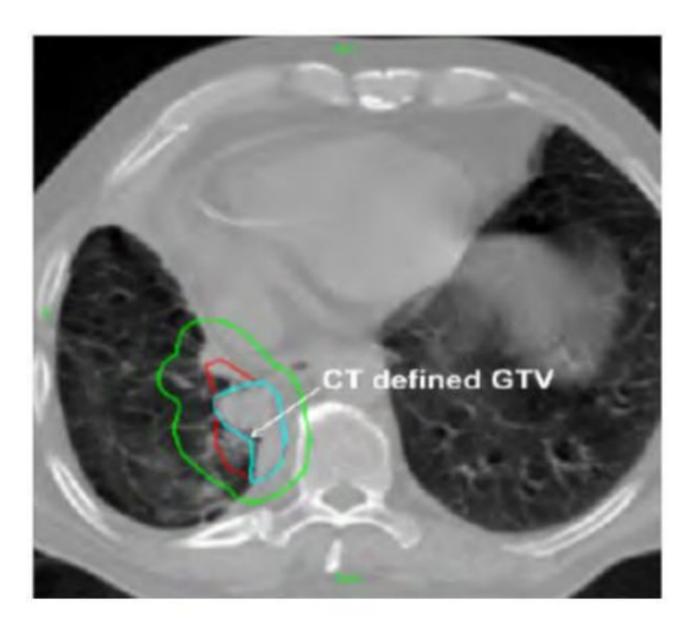
- GTV is defined based on clinical data (inspection, palpation) and imaging (CT, MR, US, PET depending on it's relevance for the tumor site)
- Definition of the GTV allows for TNM classification of the disease
- Definition of the GTV allows for tumor response assessment
- Adequate dose to GTV is therefore crucial for tumor control





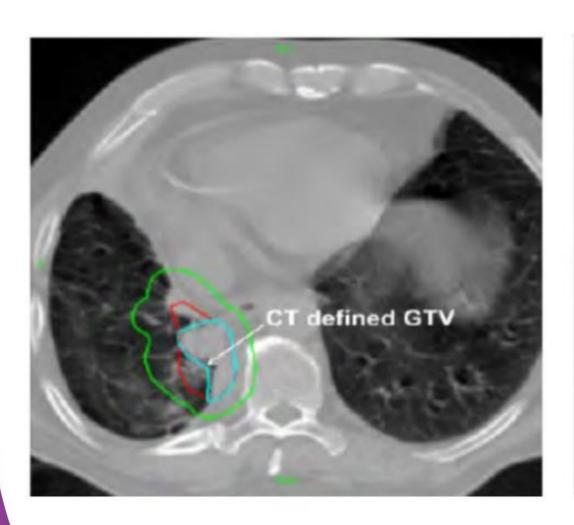


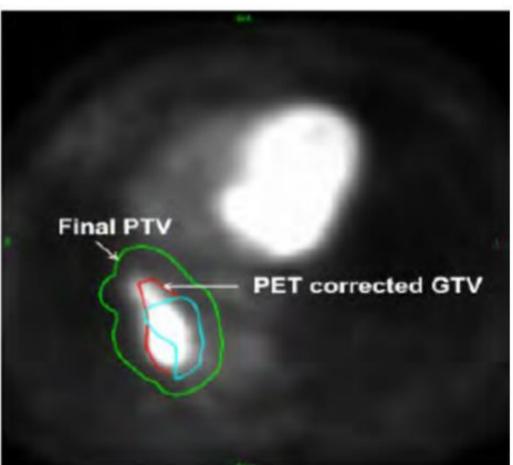
Which one is the GTV?





Are you sure about your GTV????







PET scans in delineation of lung cancer



Contents lists available at ScienceDirect



Radiotherapy and Oncology 101 (2011) 284-290



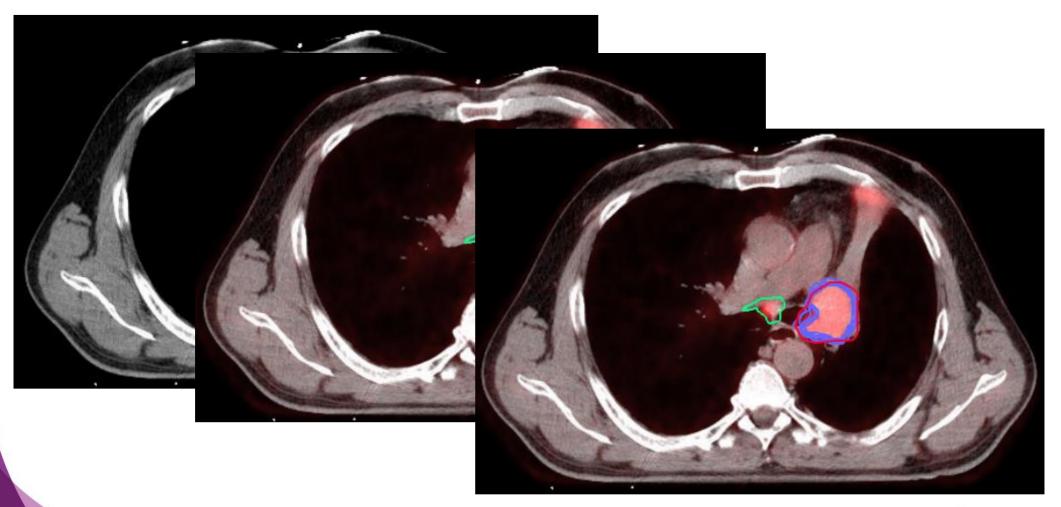
Contents lists available at ScienceDirect



- FDG-PET has an established role in contouring NSCLC
- Changes the tumor GTV in about 30–60% of patients
- Changes the nodal GTV in 9–39% of patients mainly through detection of occult metastases not seen on CT, lowering the risk of nodal recurrences



Adequate high quality imaging is a key point

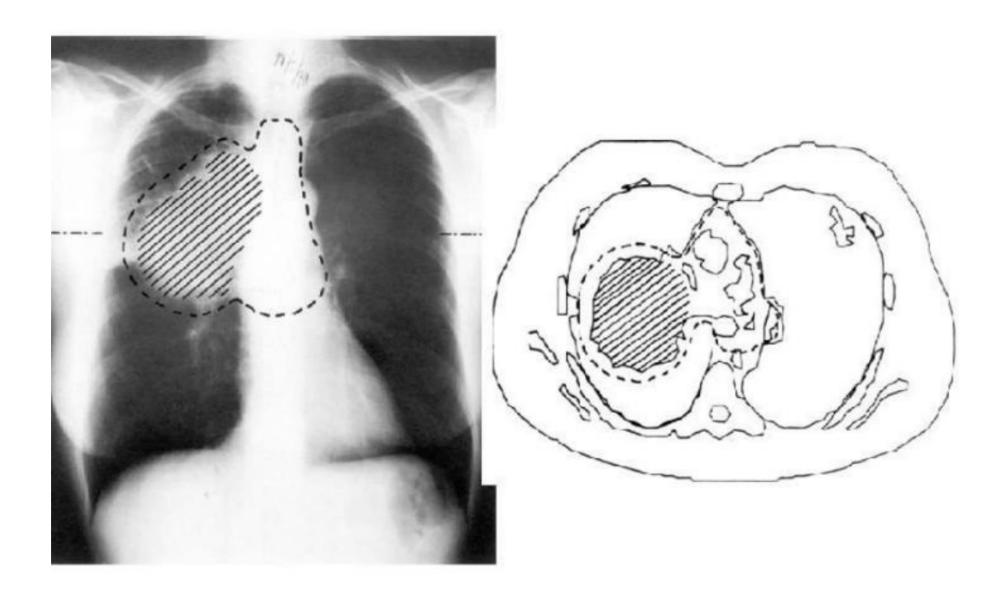


Clinical Target Volume: CTV

- Includes GTV + microscopic extension of the tumor
- Volume to adequately cover to ensure treatment efficacy weather treatment is delivered with a curative or a palliative intent
- CTV delineation is based on local and loco regional capacity/probability of extension of the tumor
- Includes potential micromets surrounding the GTV
- Includes potential micromets in tumor's drainage territory



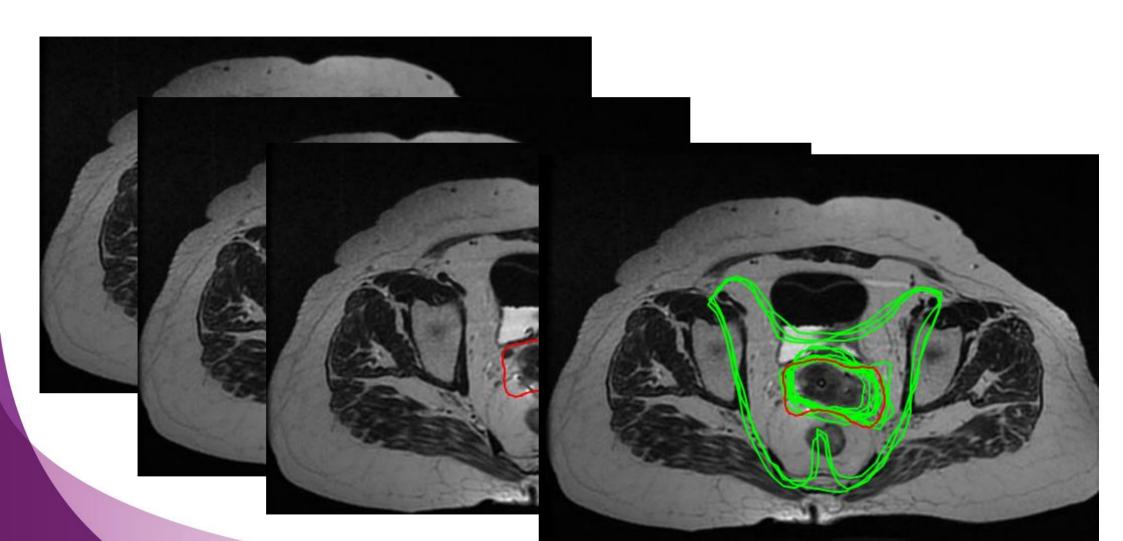
CTV





Clinical Target Volume: CTV

- High quality images are a key point for CTV delineation as well
- Margins adapted to anatomical boundaries



GTV and **CTV**

- Definition based on:
 - > Anatomy
 - Morphology
 - Imaging
 - Biology
 - Natural history of each tumor site

> But GTV and CTV delineation are independent of the technique used

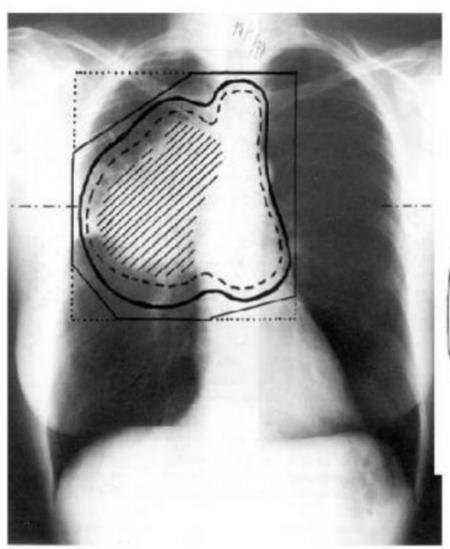


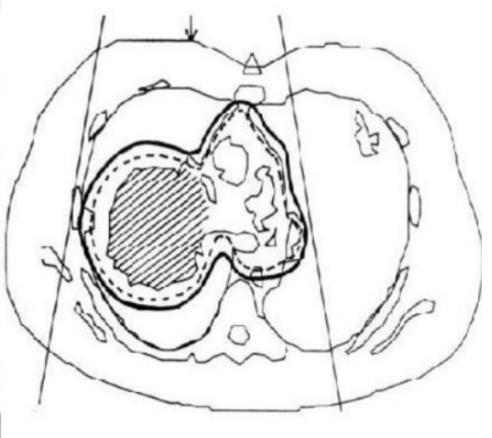
Planning Target Volume: PTV

- Geometric concept
- Meant to allow for an adequate coverage of the CTV what ever the technique, the movements, the set up uncertainties are
- Volume used for treatment planning
- Volume used for reporting



PTV





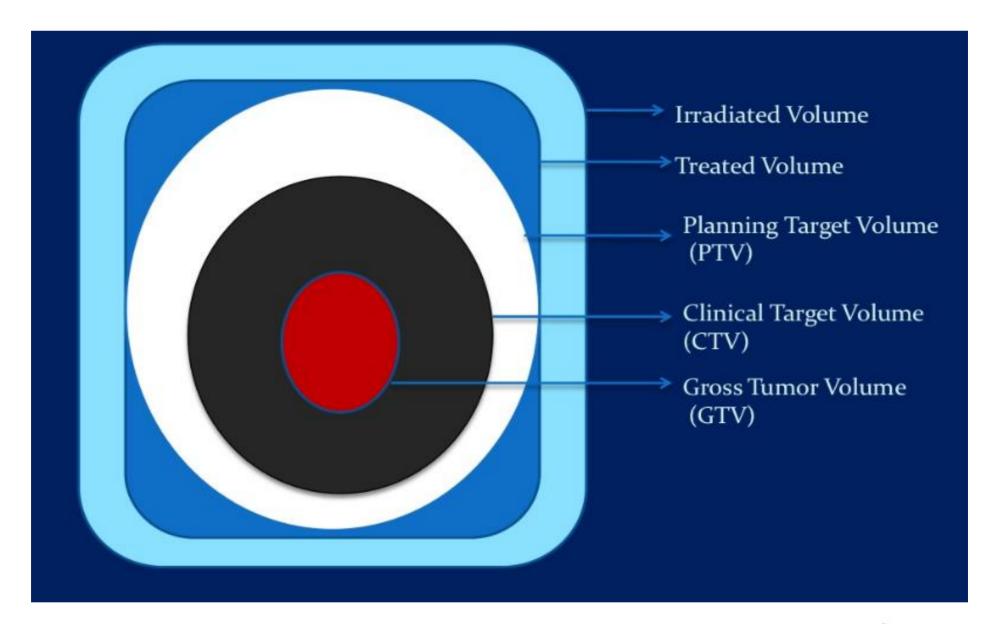


Irradiated Volume and Treated Volume: IRV and TV

- IRV: Defined as the volume receiving a significant dose on surrounding normal tissues (Organs At Risk)
- Different from the treated volume which is meant to be treated
- Both depend on the technique used
- Both can be evaluated on the dosimetry but IRV evaluation is rather limited by most TPS
 - Ex: dose estimation outside of the treated field when using non coplanar beams



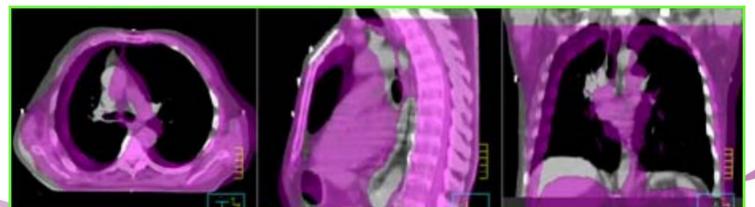
ICRU 50





ICRU 62 (in addition to ICRU 50)

- Introduces the Conformity Index: CI= treated volume/ PTV
- Recommendations on anatomical and geometrical margins
- Internal Margins: **IM** are margins integrating physiological movements (breathing, bowel/ rectum/ bladder repletion, swallowing...)
- Internal Target Volume: **ITV** is defined as the volume taking into account Internal Margins

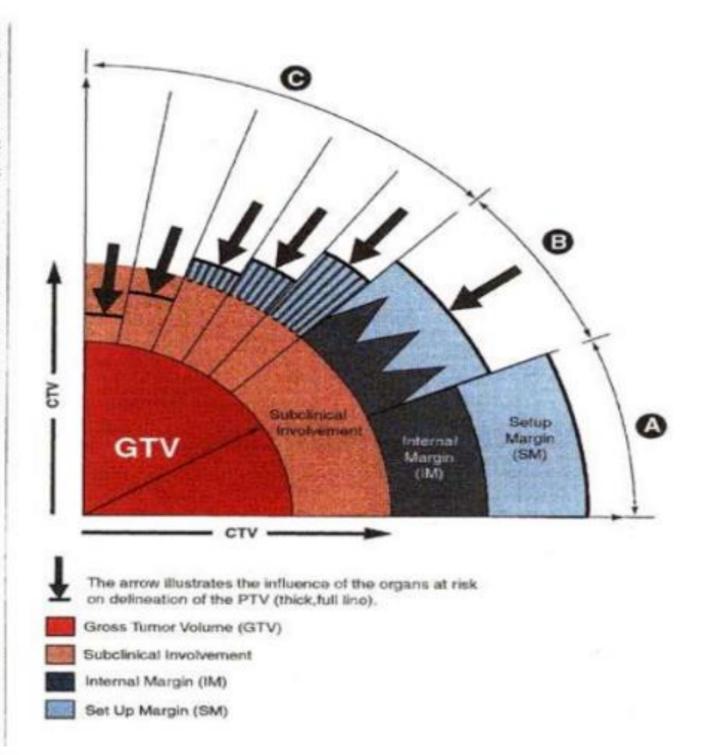




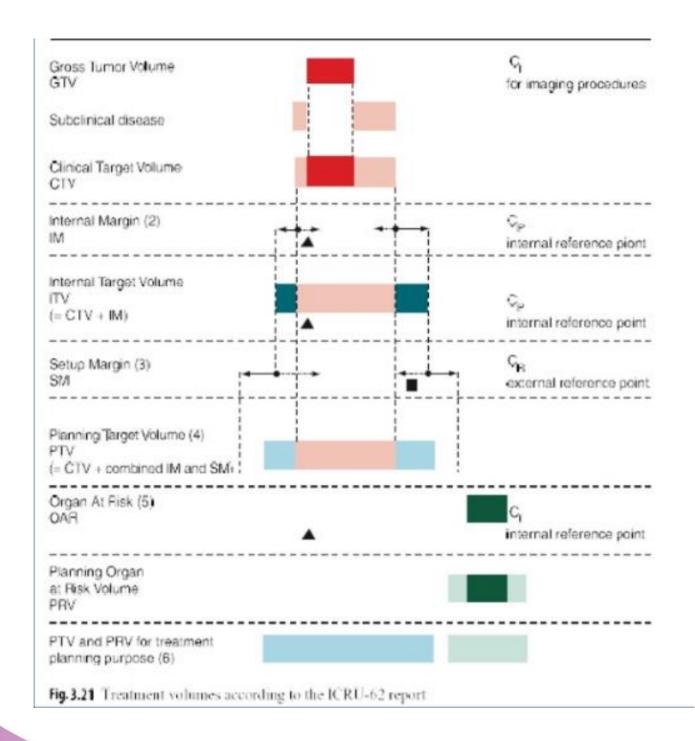
Set up Margin: SM

- Margins related to patient positioning:
 - Positioning uncertainties due to patient external movements
 - Positioning uncertainties due to body markers
 - Mechanical uncertainties due to immobilization device precision
- Depend on the technique (ex: tracking) and immobilization material and protocols (ex: thickness of painting markers or tattoos)











Contouring Guidelines

• Ex: ESTRO breast guidelines

Radiotherapy and Oncology 114 (2015) 3-10



Contents lists available at ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



ESTRO consensus guidelines

ESTRO consensus guideline on target volume delineation for elective radiation therapy of early stage breast cancer



Birgitte V. Offersen ^{a,*}, Liesbeth J. Boersma ^b, Carine Kirkove ^c, Sandra Hol ^d, Marianne C. Aznar ^e, Albert Biete Sola ^f, Youlia M. Kirova ^g, Jean-Philippe Pignol ^h, Vincent Remouchamps ⁱ, Karolien Verhoeven ^j, Caroline Weltens ^j, Meritxell Arenas ^k, Dorota Gabrys ^l, Neil Kopek ^m, Mechthild Krause ⁿ, Dan Lundstedt ^o, Tanja Marinko ^p, Angel Montero ^q, John Yarnold ^r, Philip Poortmans ^s



Contouring Guidelines

ESTRO delineation guidelines for the CTV of lymph node regions, breast and postmastectomy thoracic wall for elective irradiation in breast cancer (see figures).

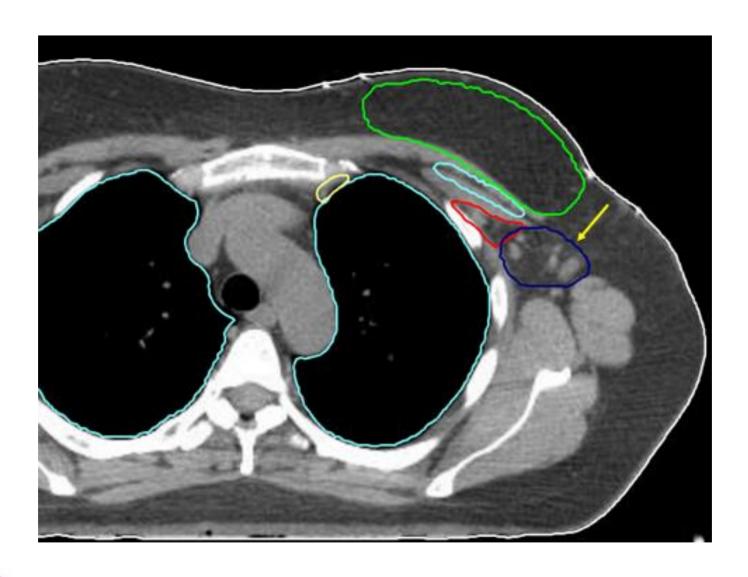
Borders per region	Axilla level 1 CTVn_L1	Axilla level 2 CTVn_L2	Axilla level 3 CTVn_L3	Lymph node level 4 CTVn_L4	Internal mammary chain CTVn_IMN	Interpectoral nodes CTVn_interpectoralis	Residual breast CTVp_breast	Thoracic wall CTVp_thoracic wall
Cranial	Medial: 5 mm cranial to the axillary vein Lateral: max up to 1 cm below the edge of the humeral head, 5 mm around the axillary vein	Includes the cranial extent of the axillary artery (i.e. 5 mm cranial of axillary vein)	Includes the cranial extent of the subclavian artery (i.e. 5 mm cranial of subclavian vein)	Includes the cranial extent of the subclavian artery (i.e. 5 mm cranial of subclavian vein)	Caudal limit of CTVn_L4	Includes the cranial extent of the axillary artery (i.e. 5 mm cranial of axillary vein)	Upper border of palpable/ visible breast tissue; maximally up to the inferior edge of the sterno- clavicular joint	Guided by palpable/visible signs; if appropriate guided by the contralateral breast; maximally up to the inferior edge of the sterno-clavicular joint
Caudal	To the level of rib 4 – 5, taking also into account the visible effects of the sentinel lymph node biopsy	The caudal border of the minor pectoral muscle. If appropriate: top of surgical ALND	5 mm caudal to the subclavian vein. If appropriate: top of surgical ALND	Includes the subclavian vein with 5 mm margin, thus connecting to the cranial border of CTVn_IMN	Cranial side of the 4th rib (in selected cases 5th rib, see text)	Level 2's caudal limit	Most caudal CT slice with visible breast	Guided by palpable/visible signs; if appropriate guided by the contralateral breast

B.Offersen et al radiother oncol 2015



Contouring Guidelines

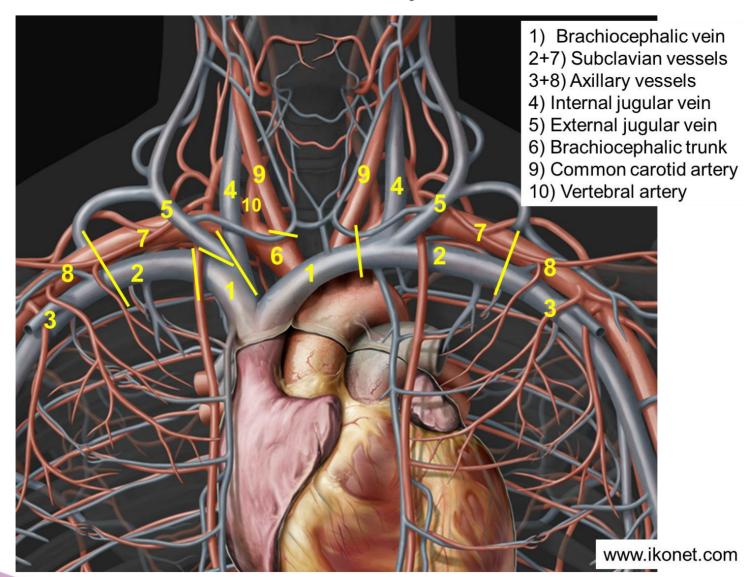
• Ex: ESTRO breast guidelines





Contouring guidelines

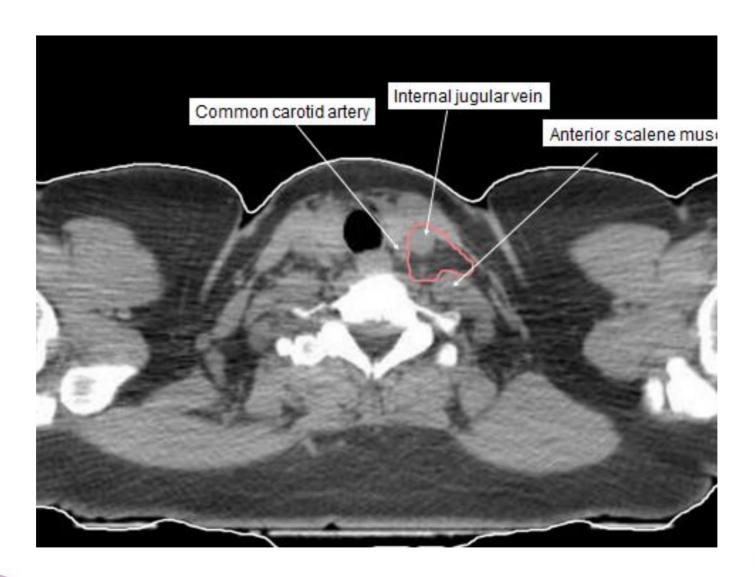
Anatomical basis are the key!





Contouring guidelines

Anatomical basis are the key!

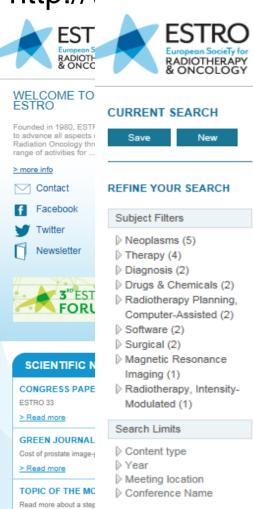




ESTRO guidelines





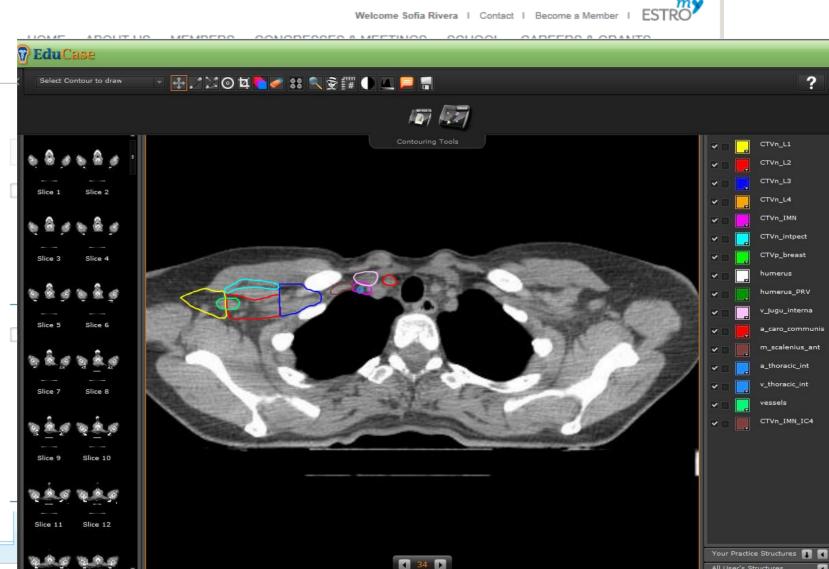


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developed by a national co

experts

> Read more



Take home messages:

- Inter observer variability in contouring can translate in a systematic error
- Need for a common language: ICRU
- Need for delineation guidelines
- Need for teaching in contouring



Thank you for you attention

Any question?



ORGANS AT RISK DELINEATION



Liz Forde, MSc (RTT)
Assistant Professor
Discipline of Radiation Therapy
Trinity College Dublin





Learning Outcomes

- Discuss the changing roles and responsibilities of RTTs for Organ at Risk (OAR) delineation
- Identify skills required to delineate OARs
- Indentify tools for implementing RTT OAR delineation into your department
- Identify common OARs based on current clinical trials and evidence based consensus guidelines
- Discuss the impact of inaccurate OAR delineation on the evaluation of plan quality

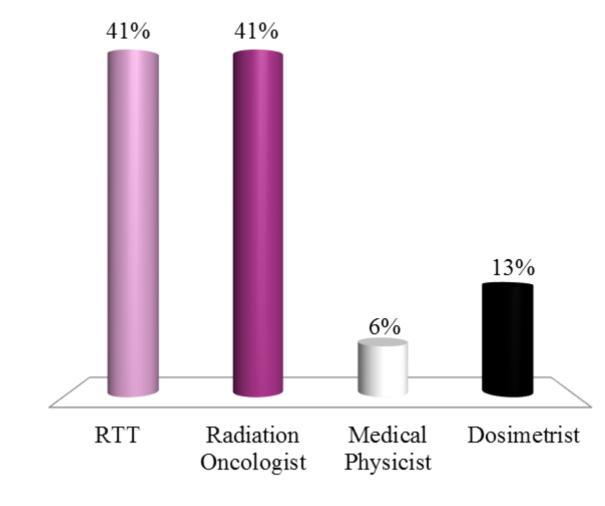
Question Time!





In my current practice organs at risk are contoured by the:

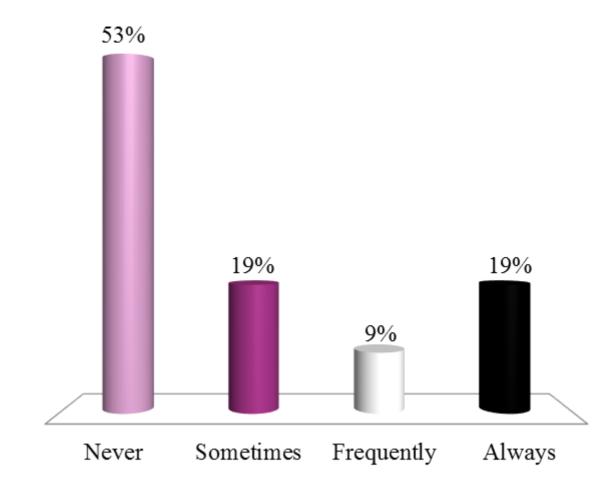
- A. RTT
- B. Radiation Oncologist
- C. Medical Physicist
- D. Dosimetrist





I personally am involved in OAR delineation:

- A. Never
- B. Sometimes
- C. Frequently
- D. Always





The New RTT!







"flexible inter professional boundaries" Schick et al., 2011







"The goal of a radiation therapist undertaking OAR delineation is logical role expansion." (Schick et al 2011)



The New RTT



Journal of Medical Imaging and Radiation Sciences

> Journal de l'imagerie médicale et des sciences de la radiation

www.elsevier.com/locate/jmir

Role Development for Radiation Therapists: An Examination of the Computed Tomographic Simulation Procedure for Patients Receiving Radiation Therapy for Breast Cancer

Journal of Medical Imaging and Radiation Sciences xx (2014) 1-8

Bonnie Bristow, MRT(T), BSc*, Saffiyya Saloojee, MRT(T), Michele Silveira, MRT(T), Shila Vakani, MRT(T) and Angela Turner, MRT(T), BA(Hons)

Department of Radiation Therapy, Odette Cancer Centre, Sunnybrook Health Sciences Centre, Toronto, Ontario, Canada

- Comparison of practice and confidence
- Identified tasks performed at CT Simulation
- Results: 84% no change made by RO

Table 4 Responses for Confidence Levels ($n=9\ ROs,\ n=21\ RTs$)

I have confidence in RTs performing the following tasks:	Agree (%)		Neutral (%)		Disagree (%)	
	RO	RT	RO	RT	RO	RT
Placement of baseline	75	100	0	0	25	0
Contouring of cardiac volume	87	95	0	5	13	0
Lung volume	76	100	12	0	12	0
Scar/seroma delineation	50	95	25	5	25	0
Cardiac contour	88	72	12	23	0	5
Spinal contour	75	90	13	10	12	0
Placement of field junction	75	80	13	10	12	10
Humeral shielding	63	62	25	24	12	14
Selection of immobilization	75	90	25	5	0	5



RTT Lead OAR Delineation – Conclusions from Literature

- Potential for site specialisation of RTs
 - Provide mentorship
 - > "train the trainers" approach

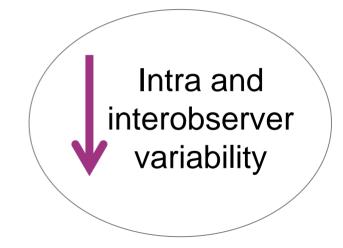
- Confidence and accuracy would improve with:
 - Standard protocols or "supporting documentation"
 - Consensus
 - Exposure to a high number of cases and "non standard" cases
 - Enhanced communication between ROs and RTs

Training model that includes case based education package and is competency based

Tools for Implementation and Facilitating Change

- Education
 - Online courses
 - > Support from national and international bodies

- Culture of the department
 - Clinical mentorship
 - Commitment to evidence based practice
 - Commitment to role development
 - Shared goals within the MDT
 - Open communication





Why Are OARs So Important?

- Do no harm culture of medicine
 - Decrease impact of radiation to our patients
- Requirement for inverse planning optimisation process
 - > IMRT
 - > VMAT
- Generates DVH information and assists in prediction of toxicity
 - > Serial and Parallel structures
 - Assessment of clinical impact and disturbance on daily activities

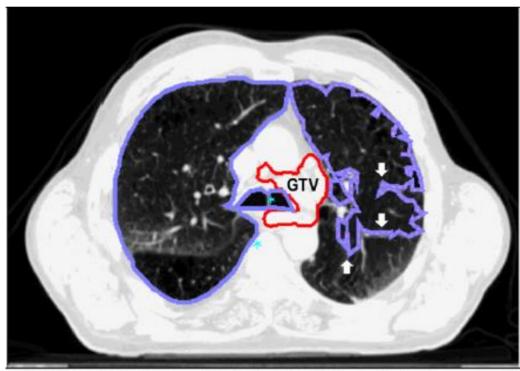


Why Is Accuracy So Important?

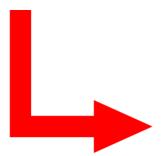
- Consistency and uniformity
 - Within the department
 - Prospective data collection
 - Analysis of local practice and impact on patients
 - Within the context of clinical trials
 - Compliance with trial specifications
 - Allows for collections of data and comparison of outcomes and toxicity at a larger international scale
 - "reduction in inter- and intra-observer variability and therefore unambiguous reporting of possible dose-volume effect relationships" (van der Water, 2009)



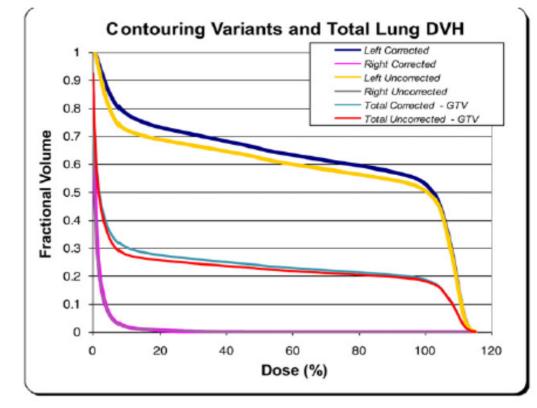
Why is *Accuracy* So Important?



A: Lung Contour - Autotrack Failure (white arrows)



What is wrong in this picture? What has caused this? What impact would this have?

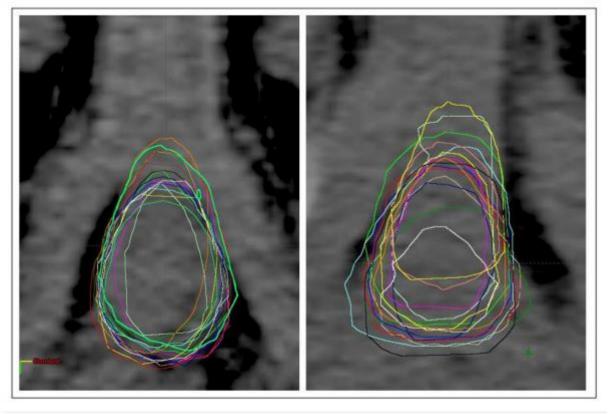


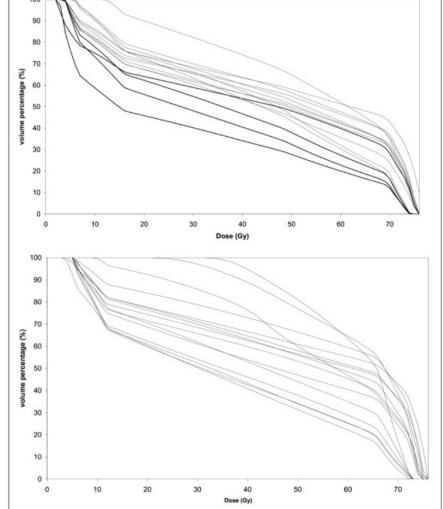
C: Lung DVH differences of contouring variants

Research Open Acc

Inter-observer variability in contouring the penile bulb on CT images for prostate cancer treatment planning

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A plot of the central slice of PB contours drawn by all observers of two patients: one with the lowest inter-observer volume variation (left side) and one with the largest interobserver volume variation (right side).

Graphs showing PB dose-volume histograms relative to the two patients in figure 5: the first (top of figure) with the lowest impact of inter-observer variation on DVH parameters, and the second (bottom of the figure) with the largest impact of inter-observer variation on DVH parameters.

Perna et al. Radiation Oncology 2011 6:123 doi:10.1186/1748-717X-6-123

Possible recommendations put forward by the authors:

Contouring by a single user Introduction of MRI into practice Improving the agreement between observers (consensus)



What Are Some of the Challenges in Delineation

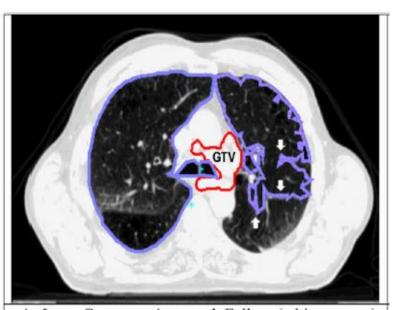
- Windowing
- Length to contour
- Over reliance on auto-contouring
- Contrast
- Motion
- Exclusion of disease
- Patient positioning



Tools Available – Auto-segmentation

- Auto segmentation based on tools in the TPS
 - Widely available
 - > Spindle snake, Flood fill...

Common errors include...using the auto-threshold contouring tools in the TPS and not editing the resulting errors" (*Gay et al., 2012*)



A: Lung Contour - Autotrack Failure (white arrows)

Trachea included and portion of lung missing



Tools Available – Auto-segmentation

- Atlas based Auto segmentation
- Propagation of segmented structures from an atlas onto the patient image using deformable registration (Lim and Leech, 2017)
- Atlas can be based on:
 - Single patient dataset
 - Multiple patient data (based on an average of a range of patients from multiple libraries)
 - Model based (using library of previously manually contoured patients)



Tools Available – Auto-segmentation

- Attractive due to time saving aspects and support of adaptive RT, but...
- Beware of automaticity!
 - Even with the implementation of AS software in the future, it should be reinforced that manual editing is still a necessity for patient safety." (*Lim and Leech, 2017*)
 - "atlas-based automatic segmentation tool ... is timesaving but still necessitates review and corrections by an expert" (Daisne and Blumhofer, 2013)

You still need the anatomical knowledge of what is correct!



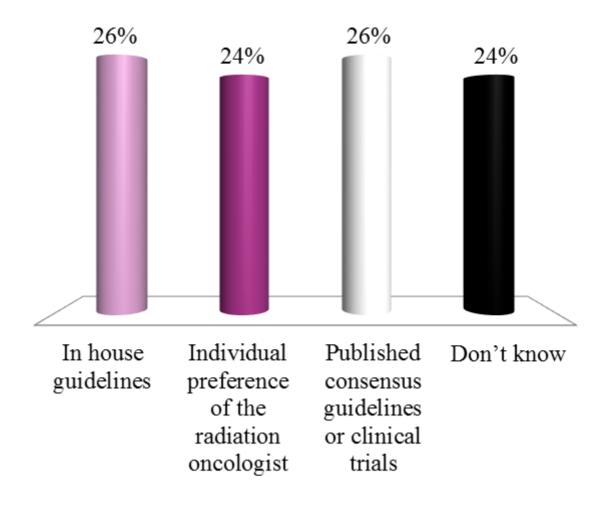
Question Time!





In your current practice what defines how organs at risk are contoured?

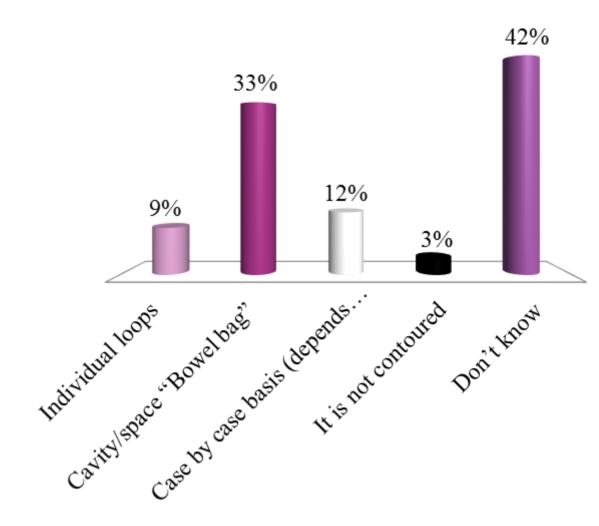
- A. In house guidelines
- B. Individual preference of the radiation oncologist
- C. Published consensus guidelines or clinical trials
- D. Don't know





In your current practice how is the small bowel contoured?

- A. Individual loops
- B. Cavity/space "Bowel bag"
- C. Case by case basis (depends on treatment site)
- D. It is not contoured
- E. Don't know





Is there Consensus?

QUANTEC

Contouring Atlases eLearning
Modules by
Experts

Clinical Trials



Let's Look at Some Common OARs in the

Pelvis

Rectum

Small Bowel





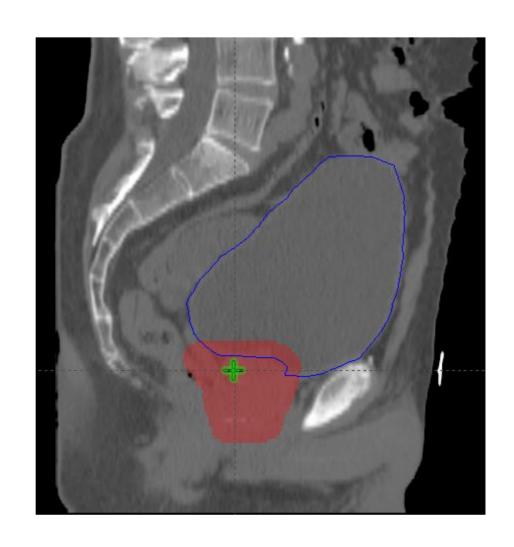
Bladder

Sigmoid

Femoral heads



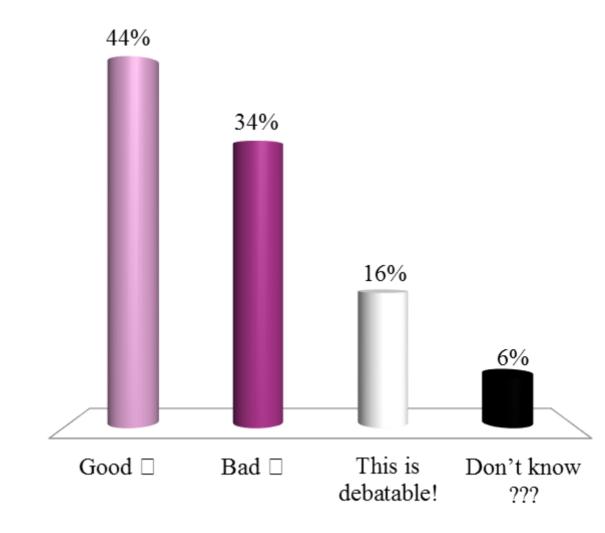
Bladder - Good or Bad?





This bladder size is:

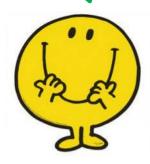
- A. Good ©
- B. Bad 🕾
- C. This is debatable!
- D. Don't know???



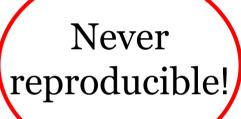


Bladder - Good or Bad?

Fantastic DVH!











What Do the Experts Say? - Bladder

- Uncertainties or variations in practice:
 - ➤ Bladder wall or solid contour including urine?
 - ➤ Whole structure or set length from PTV?
 - Contrast from post prostatectomy (defining the SUA)
 - Easy to define on planning CT but potential of high variation
 - Unrealistic DVH
 - Consider CBCT review and generate bladder DVH of the day
 - Does it impact on target position?
 - What are you treating?
 - Prostate
 - Prostate bed
 - Endometrial cancer



What Do the Experts Say? - Rectum

- Uncertainties or variations in practice:
 - ➤ Inferior limit Anal verge or ischial tuberosities?
 - Rectal wall or solid including contents?
 - > Set length defined by the PTV volume?
- Recommendations:

Organ segmentation

The rectum should be segmented from above the anal verge to the turn into the sigmoid colon, including the rectal contents. Although there can be variation in defining these landmarks, the superior limit is where the bowel moves anteriorly, close to the inferior level of the sacroiliac joints, and the inferior limit is commonly at the bottom of the ischial tuberosities. In prostate cancer therapy, an empty rectum at





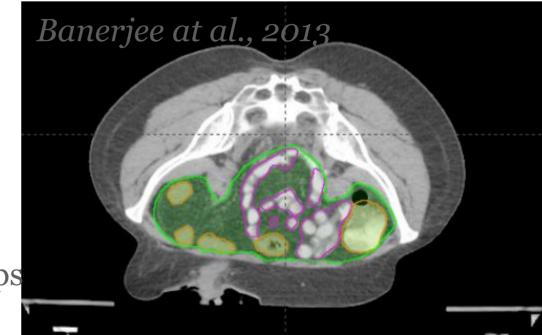
What Do the Experts Say? – Small Bowel

- Uncertainties or variations in practice
 - ➤ What is large bowel/vessels/nodes
 - Oral contrast results in artefact on planning scan and inappropriate
 HU
 - > Small bowel position is variable during treatment
 - Individual loops vs. "Bowel bag"

• Recommendations:

The absolute volume of small bowel receiving ≥15 Gy should be held to <120 cc when possible to minimize severe acute toxicity, if delineating the contours of bowel loops themselves. Alternatively, if the entire volume of peritoneal space in which the small bowel can move is delineated, the volume receiving >45 Gy should be <195 cc when possible.

Orange = Large bowel
Pink = Small bowel loops
Green = Bowel bag









Atlases available online at: www.rtog.org/CoreLab/ContouringAtlases.aspx

Int J Radiation Oncol Biol Phys. 2012; 83(3):

Clinical Investigation: Genitourinary Cancer 353-362 ■ PenileBulb Bladder ■ Prostate Femur L Femur R BowelBag Coronal Femur Rand Femur Linclude

PenileBulb

SmallBowel

SeminalVeso

AnoRectumS

Bladder

Prostate

PenileBulb has a

rounded shape

the proximal femur, not just the ball of the femur

Pelvic Normal Tissue Contouring Guidelines for Radiation Therapy: A Radiation Therapy Oncology Group Consensus **Panel Atlas**

Hiram A. Gay, M.D.,* H. Joseph Barthold, M.D.,†,‡ Elizabeth O'Meara, C.M.D.,§ Walter R. Bosch, D.Sc.,* Issam El Naga, Ph.D., Rawan Al-Lozi, B.A.,* Seth A. Rosenthal, M.D., Colleen Lawton, M.D., ** W. Robert Lee, M.D., Howard Sandler, M.D., ** Anthony Zietman, M.D., ** Robert Myerson, M.D., Ph.D., * Laura A. Dawson, M.D., Christopher Willett, M.D., Lisa A. Kachnic, M.D., Anuja Jhingran, M.D., *** Lorraine Portelance, M.D., ††† Janice Ryu, M.D., William Small, Jr., M.D., Htt David Gaffney, M.D., Ph.D., SSS Akila N. Viswanathan, M.D., M.P.H., and Jeff M. Michalski, M.D.*

Sagittal

Contour BowelBag, Colon and SmallBowel the

recommended cm above PTV not necessarily this high

Any sigmoid adjacent or above the uterus or a brachytherapy applicator should be contoured.

Sigmoid

AnoRectum

BowelBag UteroCervix

Bladder

Let's Look at Some Common OARs in the

Thorax

Heart

Lungs



Ribs

Spinal Cord

Oesophagus

Main Bronchus

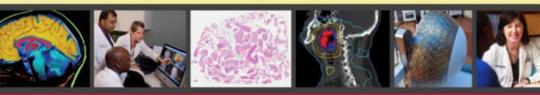
Brachial Plexus



RTOG Thoracic Atlas available from:

http://www.rtog.org/CoreLab/ContouringAtlases/LungAtlas.aspx





Atlases for Organs at Risk (OARs) in **Thoracic Radiation Therapy**

Feng-Ming (Spring) k **Leslie Quint** Mitchell Machi **Jeffrey Bradk**



Int. J. Radiation Oncology Biol. Phys., Vol. 81, No. 5, pp. 1442-1457, 2011 Copyright © 2011 Elsevier Inc. Printed in the USA. All rights reserved 0360-3016/\$ - see front matter

doi:10.1016/j.ijrobp.2010.07.1977

CLINICAL INVESTIGATION

Normal Tissue

CONSIDERATION OF DOSE LIMITS FOR ORGANS AT RISK OF THORACIC RADIOTHERAPY: ATLAS FOR LUNG. PROXIMAL BRONCHIAL TREE. ESOPHAGUS. SPINAL CORD, RIBS, AND BRACHIAL PLEXUS

FENG-MING (SPRING) KONG, M.D., Ph.D.,* TIMOTHY RITTER, Ph.D.,* DOUGLAS J. QUINT, M.D., SURESH SENAN, M.D., LAURIE E. GASPAR, M.D., RITSUKO U. KOMAKI, M.D.,

COEN JEFFREY D. BRA



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0360-3016/\$-see front matter

doi:10.1016/j.jjrobp.2009.10.058

CLINICAL INVESTIGATION

Breast

DEVELOPMENT AND VALIDATION OF A HEART ATLAS TO STUDY CARDIAC EXPOSURE TO RADIATION FOLLOWING TREATMENT FOR BREAST CANCER

MARY FENG, M.D.,* JEAN M. MORAN, Ph.D.,* TODD KOELLING, M.D.,[†] AAMER CHUGHTAI, M.D.,[‡] JUNE L. CHAN, M.D., * LAURA FREEDMAN, M.D., * JAMES A. HAYMAN, M.D., * RESHMA JAGSI, M.D., D. PHIL., * SHRUTI JOLLY, M.D., * JANICE LAROUERE, M.D., * JULIE SORIANO, M.D.,* ROBIN MARSH, C.M.D.,* AND LORI J. PIERCE, M.D.*

What Do the Experts Say? - Lung

Challenges

- Inappropriate window settings!
- Exclusion of disease from healthy lung?
- Inclusion of vessels?

Recommendations

- Air inflated lung only
 - Do not include fluid
- Contoured as single or combined structures
- Exclude lung GTV
- Exclude trachea/bronchus
- Exclude vessels <1cm
- Auto-segmentation is allowed combined with manual inspection
- Ensure appropriate windowing



What Do the Experts Say? – Spinal Cord

Challenges

- Difficult to see true cord on CT
- Often not specifically covered in atlases
- Circumferential extend?
 - Contour cord or canal?
- Superior/Inferior extent
 - Entire length visible on planning scan or set distance from PTV?

Recommendations

- Use MRI fusion, if available
- Contour to the bony limits of the canal
- For lung cases, superior limit is the same as oesophagus (cricoid cartilage)
- Inferior limit is L2/L3 junction



What Do the Experts Say? – *Heart*

Challenges

- Contour specific structures within the heart?
- Superior limit

Recommendations

- Superiorly: Just inferior to the left pulmonary artery, include the great vessels in a rounded contour
- Inferiorly: to diaphragm, include pericardium
- If contrast is used, contour SVC separately

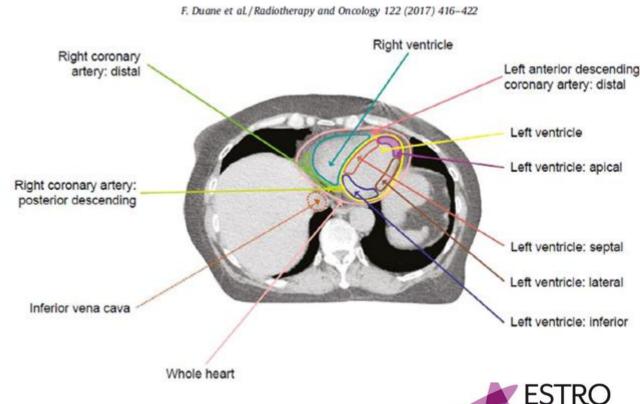


What Do the Experts Say? - Heart (Substructures)

2017 Atlas in Green Journal

 Whole heart dose may not be the best predictor for the different types of radiation induced cardiac toxicity

Focus on left ventricle and coronary arteries



Dunane et al., 201

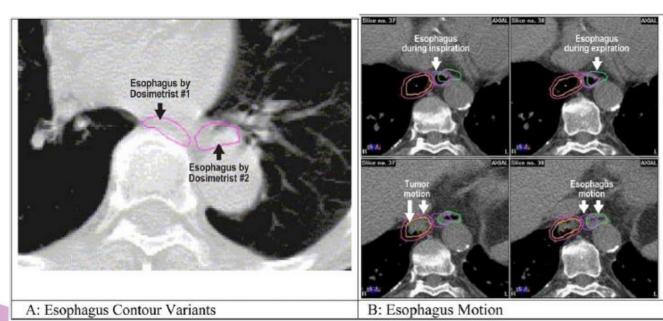
What Do the Experts Say? – *Oesophagus*

Challenges

- Impact of windowing
- Impact of oral contrast
- Motion
- Inclusion of the muscular wall
- Length of contour

Recommendations

- Use mediastinal windowing level
- Contour from cricoid cartilage to gastro oesophageal junction
- Avoid oral contrast
 - Distorts shape and density





What about clinical trials?





RADIATION THERAPY ONCOLOGY GROUP

RTOG 0529

A Phase II Evaluation of Dose-Painted IMRT in Combination with 5-Fluorouracil and Mitomycin-C for Reduction of Acute Morbidity in Carcinoma of the Anal Canal

<u>Critical Normal Structures</u>: In addition, surrounding critical normal structures, including the femoral heads (right and left), bladder, external genitalia, iliac crest, small bowel, large bowel outside the CTVs, and perianal skin should be outlined. The normal tissues will be contoured and considered as solid organs. The tissue within the skin surface and outside all other critical normal structures and PTVs is designated as unspecified tissue.

Critical normal structures: DVHs must be generated for all critical normal structures.

NOTE: Effort should be made to achieve the listed dose constraints to normal tissues below. Failure to meet the 6.5.1.1 and 6.5.1.2 dose constraints will result in minor deviation. The dose constraints are listed in order from most to least important.



www.rediournal.org

AGITG – For Anus

Bladder

- Entire outer wall
- Femoral Heads
 - ➤ Inferior Cranial edge of the lesser trochanter
- Bowel
 - Small and large bowel
 - > 15mm superior of PTV down to the rectosigmoid junction
- External Genitalia
 - ➤ Male penis, scrotum, skin and fat anterior to the pubic symphysis
 - Female clitoris, labia majora and minora, skin and fat anterior to pubic symphysis
- Bone Marrow
 - Iliac crests, both contoured and combined
 - Superior top of the iliac crests
 - Inferior superior part of the acetabulum

Clinical Investigation: Gastrointestinal Cancer

Australasian Gastrointestinal Trials Group (AGITG) Contouring Atlas and Planning Guidelines for Intensity-Modulated Radiotherapy in Anal Cancer

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Sarat Chander, M.B.B.S., F.R.A.N.Z.C.R.,† Julie Chu, M.B.B.S., F.R.A.N.Z.C.R.,†
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RAVES



• Femoral head:

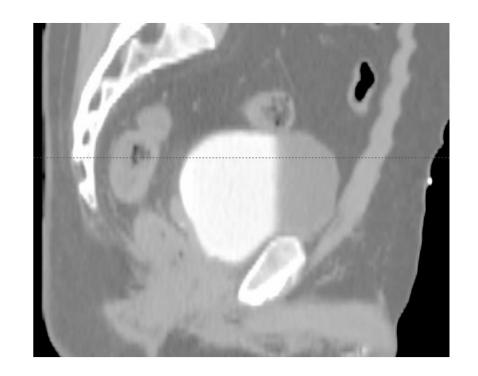
- Superior acetabulum
- ➤ Inferior inferior edge of the treatment field

• Bladder:

Whole structure with bulk homogeneity correction for contrast

• Rectum:

- Superior rectosigmoid junction
- ➤ Interior 15mm inferior to the CTV



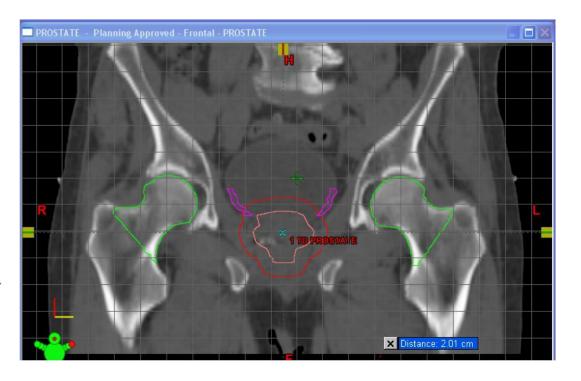


PROFIT Trial

Rectal Wall

Bladder Wall

 Femoral Head and Neck





RADIATION THERAPY ONCOLOGY GROUP

RTOG 0615

A PHASE II STUDY OF CONCURRENT CHEMORADIOTHERAPY USING THREE-DIMENSIONAL CONFORMAL RADIOTHERAPY (3D-CRT) or INTENSITY-MODULATED RADIATION THERAPY (IMRT) + BEVACIZUMAB (BV) FOR LOCALLY OR REGIONALLY ADVANCED NASOPHARYNGEAL CANCER

NCI-supplied agent: Bevacizumab (NSC 704865; IND 79211)

A lot of contouring!

Critical Normal Structures

Surrounding critical normal structures, including the brainstem, spinal cord, optic nerves, chiasm, parotid glands, pituitary, temporo-mandibular (T-M) joints and middle and inner ears, skin (in the region of the target volumes), oral cavity, mandible, eyes, lens, temporal lobes, brachial plexus, esophagus (including postcricoid pharynx) and glottic larynx should be outlined

Planning Priorities

Critical structures are

critical!

Critical normal structure constraints followed by the prescription goals are the most important planning priorities. The priorities in addressing the protocol aims and constraints will be in the following order:

- Critical Normal Structure Constraints (Section 6.5);
- 2) Dose Specifications (Section 6.1);
- Planning Goals: Salivary glands (Section 6.5.3);
- 4) Planning Goals: Other normal structures (Section 6.5.3).



• RTOG Atlases for H&N do not cover OARs!!!



- Where to turn to?
 - Published literature
 - Expert consensus



Radiotherapy and Oncology 117 (2015) 83-90



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Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



Head and neck guidelines

CT-based delineation of organs at risk in the head and neck region: DAHANCA, EORTC, GORTEC, HKNPCSG, NCIC CTG, NCRI, NRG Oncology and TROG consensus guidelines



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Consensus panel of Radiation Oncologists from Europe, North America, Asia and Australia



• Don't worry – even the "experts" have significant interobserver variability

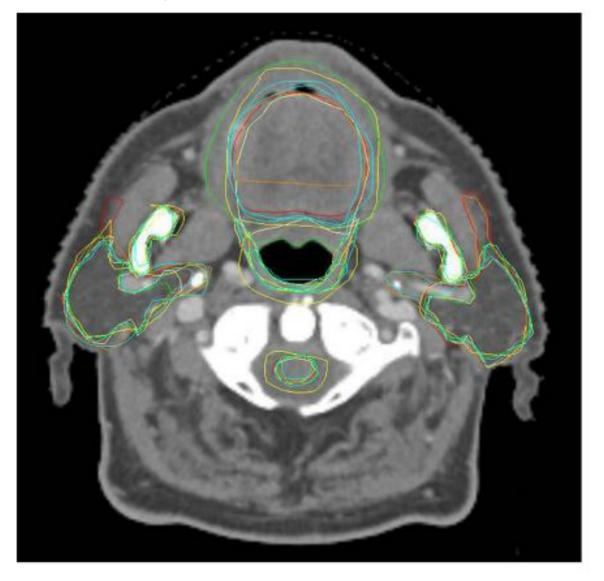


Fig. 1. Delineation results of 7 members of the panel for the parotid glands, spinal cord, pharyngeal constrictor muscles and the oral cavity, projected on an axial CT slice.

- But still worth a read!
- Text and table description of anatomy with multimodality images to show

Supraglottic larynx

The supraglottic larynx is delineated according to Christianen et al. [7]. Anatomic borders are listed in Table 1. An axial slice of the supraglottic larynx is depicted in Fig. 4a.

Glottic area

We decided to define the glottic area structure, including the vocal cords and paraglottic fat. Air should be excluded from the contour. Cranial, caudal and posterior borders can be found in Table 1. An axial slice of the glottic area is depicted in Fig. 4b.

Arytenoids

The arytenoids (or arytenoids cartilage) are defined as a separate structure. The base (caudal edge) of each arytenoid is broad for articulation with the cricoid cartilage. The apex (cranial edge) is pointed.

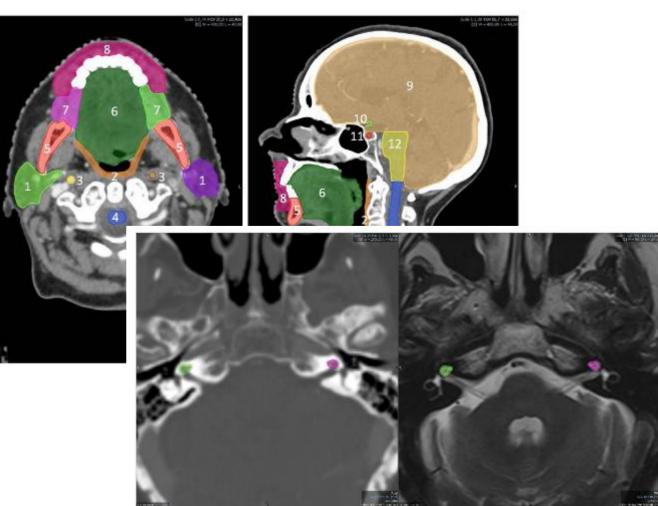
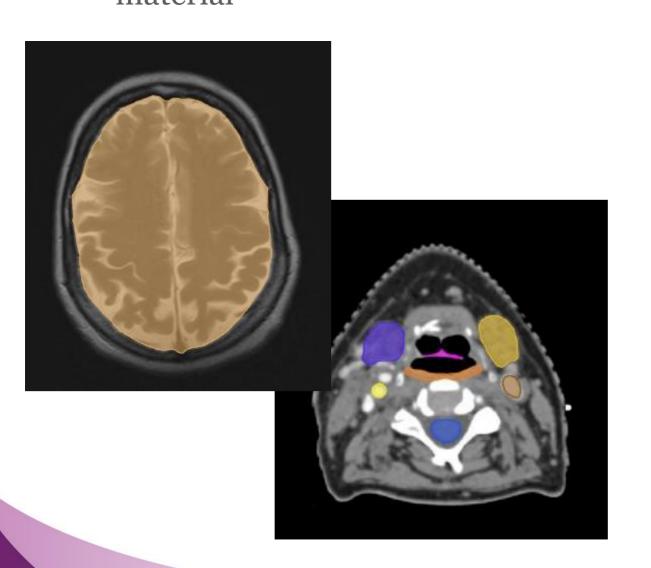


Fig. 2. Delineation of the cochlea in CT bone settings (left), matched to MRI-T2 (right).

 Thank you – they have an atlas published as supplementary material



Anterior segment of the eyeball L Anterior segment of the eyeball R Posterior segment of the eyeball L Posterior segment of the eyeball R Lacrimal gland L Lacrimal gland R Parotid gland L Parotid gland R Submandibular gland L Submandibular gland R Extended oral cavity Lips Mandible Cochlea L Cochlea R Pharyngeal constrictor muscles Glottic area Spinal cord Brachial plexus L Carotid artery L Brachial plexus R Carotid artery R Thyroid gland Buccal mucosa R Brain Buccal mucosa L Brainstem Arytenoid L Pituitary gland Arytenoid R Optic chiasm Crico-pharyngeal inle Optic nerve L Cervical esophagus

Supraglottic larynx

Optic nerve R





Radiation Oncology, Head and Neck, Organs at Risk (OAR)

ID: 001523 Approved:08 Aug 2013 Last Modified: 02 Oct 2013 Review Due:08 Se

Head and Neck Organs At Risk (OAR)

Doses listed in the table below are based on radiation doses of 2Gy per fraction

OAR Structure	Description based on RTOG 0920	True structure constraint (ideal)	Notes (Aim to keep doses as low as possible)
Brainstem	The inferior most portion of the brainstem is at the cranial-cervical junction where it meets the spinal cord. The superior most portion of the brainstem is approximately at the level of the top of the posterior clinoid. The brainstem shall be defined based on the treatment planning CT scan.	■ Max dose ≤ 54Gy	Additional goals may include: ■ ≤ 1% of PRV to exceed 60Gy ■ small volumes (1-10cc) max dose ≤ 59Gy for fraction doses ≤ 2Gy ¹
Optic nerves		■ Max dose ≤ 50Gy	Additional goals may include: ■ ≤ 1% of PRV to exceed 60Gy ■ To keep the risk of radiation induced optic neuropathy (RION) ≤ 3-7%, max dose 55-60Gy ■ The risk of RION increases to 7-20% for doses > 60Gy in 1.8-2Gy fractions ²
Optic Chiasm		■ Max dose ≤ 54Gy	Additional goals may include: ■ ≤ 1% of PRV to exceed 60Gy ■ To keep risk of radiation induced optic neuropathy (RION) < 3-7%, max dose 55-60Gy ■ The risk of RION increase to 7-20% for doses > 60Gy in 1.8-2Gy fractions ²



eviQ Head and Neck Critical Structures Atlas

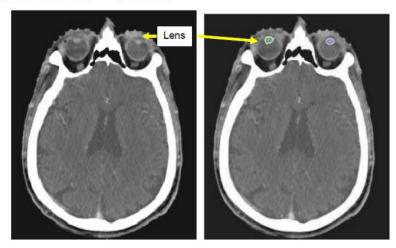


LENSES

Description: The lens is an anterior structure 5-8mm in length

Window level: W600/L40

Typical tolerance dose: 5Gy maximum



LACRIMAL GLAND

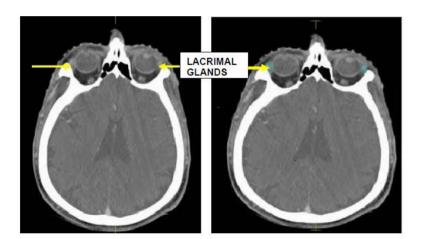
Location: Bilateral structure located supero-laterally to the orbits.

Length: Contoured extends approximately 10-15mm

Scroll through the images first to distinguish between muscle and the gland itself.

Window level: use approximately W270/L40 or W500/L60.

Typical Tolerance dose: <40Gy

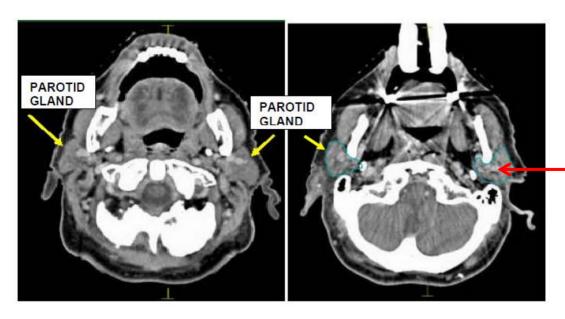


- Shows adjacent images with and without contour
- Provides anatomical location, description, suggested window level and tolerance dose





eviQ Head and Neck Critical Structures Atlas



Note: degradation of image quality due to dental artefact

PAROTID GLAND

Location: The parotid gland is a bilateral glandular tissue lying anterior to the ear between the masseter muscle and the skin. It lies inferior to the zygomatic arch beneath the skin that covers the lateral and posterior surface of the mandible.

Length: approximately 50-60mm.

Borders: Medial Border is at the styloid process. Anteriorly hooks around the posterior aspect of anterior ramus of mandible.

Window level: use approximate window levels W290/L40 or W400/L80.

Typical tolerance doses: Mean parotid dose <26Gy (in at least one gland) or at least 50 % of one gland should receive <30Gy mean. (Doses to the parotid should be kept as low as reasonably achievable)





eviQ Head and Neck Critical Structures Atlas

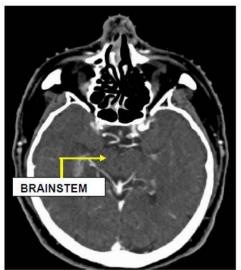
BRAINSTEM

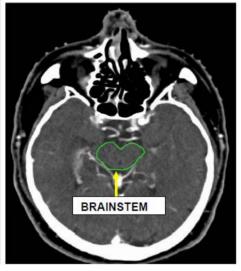
Location: Superior to as well as a continuous structure from the proximal spinal cord, the brain stem is characterised by a sudden increase in width compared to the spinal cord above the level of the foramen magnum. The brain stem sits inferior to the optic chiasm. The brainstem is composed of the medulla, the pons and midbrain. It lies posterior to the bony clivus. Use sagittal view for ant/post definition; inferiorly it may be over contoured as spinal cord, due to tighter dose constraints.

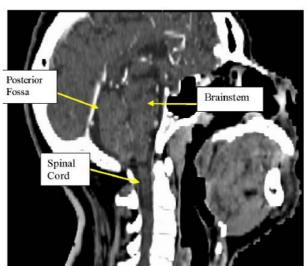
Window level: Approximate window level of W200/L80.

Typical tolerance doses: <54Gy maximum.

Remember to view structures in all planes









Superior level

Note: It is important to avoid a gap between the brainstem and spinal cord contours as the emetic centre which controls nausea is located in this space. To avoid dose dumping in this region it is advised to overlap the contours by 1 CT slice to avoid a gap between these structures. (Monroe et al 2008 Radiother Oncol 87(2):188-194).





eviQ Head and Neck Critical Structures Atlas

OPTIC CHIASM

Location: A butterfly-shaped structure which sits directly above the pituitary fossa. To aid in contouring trace the optic nerves to the point of posterior intersection to help with identifying the optic chiasm.

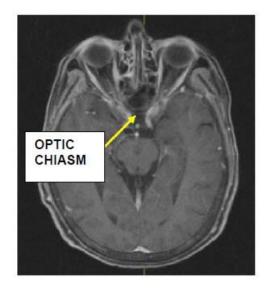
Course: Anterior it begins directly posterior to the optic canal. Begin contouring at this position. At this area it lies medial to the carotid arteries (which enhance with contrast) but anterior to the pituitary stalk. The optic chiasm joins in front of the pituitary stalk and then divides again posteriorly to travel to the most superior/anterior part of the brainstem (ie gives position of the most superior limit of the brainstem). It should look butterfly shaped.

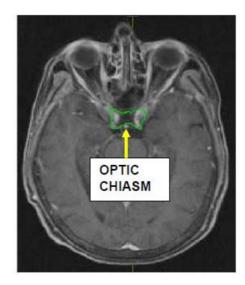
Length: Approximately 5-8mm, consider using an MRI study set if it is available for easier visualisation of the optic chiasm.

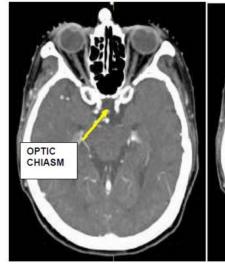
Window level: use approximate window level of W220/L70.

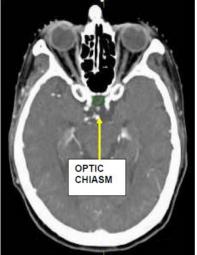
Typical Tolerance dose: Ideally <50Gy with a maximum of <54Gy

Remember to use all imaging available for that patient







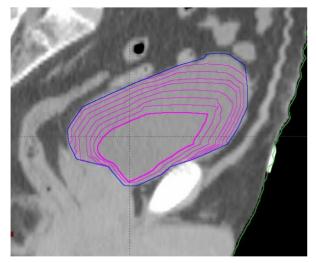




Other Points to Consider

- Planning Risk Volume
 - Margin added to true structure
 - > ICRU 83
 - > RTOG H&N Trials

Additional structures to assist in decision making at the linac



Moore and Forde, JMIRS 2017

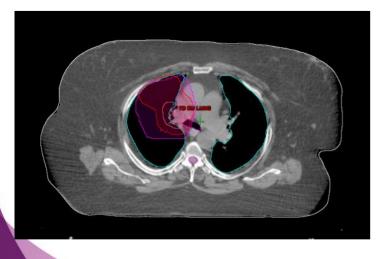
Minimum allowable bladder volume



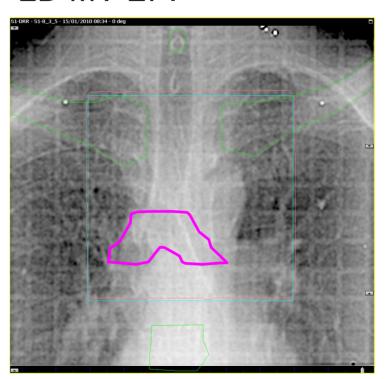
Other Structures for IGRT at the Linac

- What is the best surrogate for the target?
- What else can you see that might help you match?

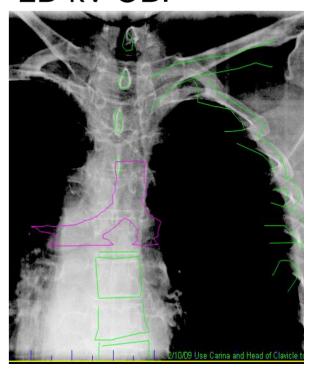
Planning CT



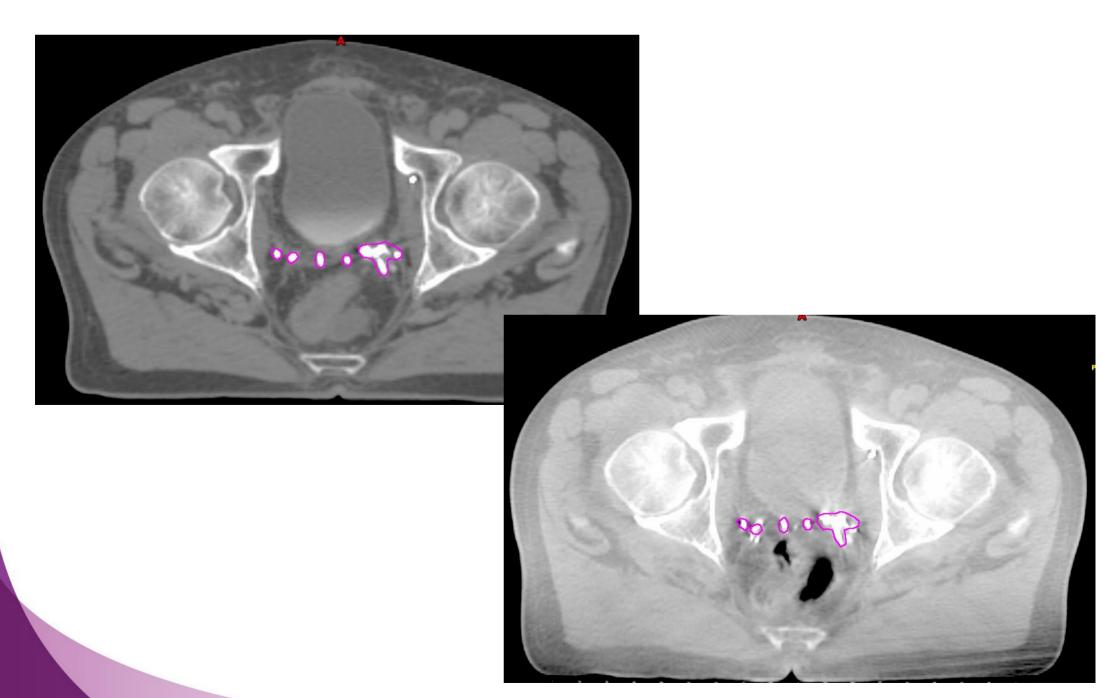
2D MV EPI



2D kV OBI



Other Structures for IGRT at the Linac



Take Home Message

- Quality assurance of organ delineation is vital regardless of who is responsible for OAR delineation
- "The accuracy of any autosegmenting tools should be carefully assessed" (Marks et al., 2010)
- Use all imaging modalities and viewing planes that you have available for that patient
- Think about the whole patient pathway
 - What will these contours impact on?
- Be consistent!
 - Preferably with international recommendations/consensus
 - At least at a local level



"Inaccuracy and variation in defining critical volumes will affect everything downstream: treatment planning, dose—volume histogram analysis, and contour based visual guidance used in image-guided radiation therapy"

(Nelms et al., 2012)



OAR delineation workshop



Sofia Rivera, M.D.
Radiation Oncology Department
Gustave Roussy
Villejuif, France



Advanced skills in modern radiotherapy
June 2017

ESTRO VISION 2020

Every cancer patient in Europe will have access to state of the art radiation therapy, as part of a multidisciplinary approach where treatment is individualized for the specific patient's cancer, taking into account the patient's personal circumstances

Radiotherapy & Oncology 103(2012) 99-101

FALCON Vison

- * Falcon provides valuable global educational training for radiotherapy professionals on delineation.
- ❖ It is a dynamic and evolving pedagogical programme that answers educational needs of current and future clinical practice using live, online and blended format.
- * It facilitates and serves research projects and development of guidelines.
- * It is fully integrated into the ESTRO School's programme and strategy.

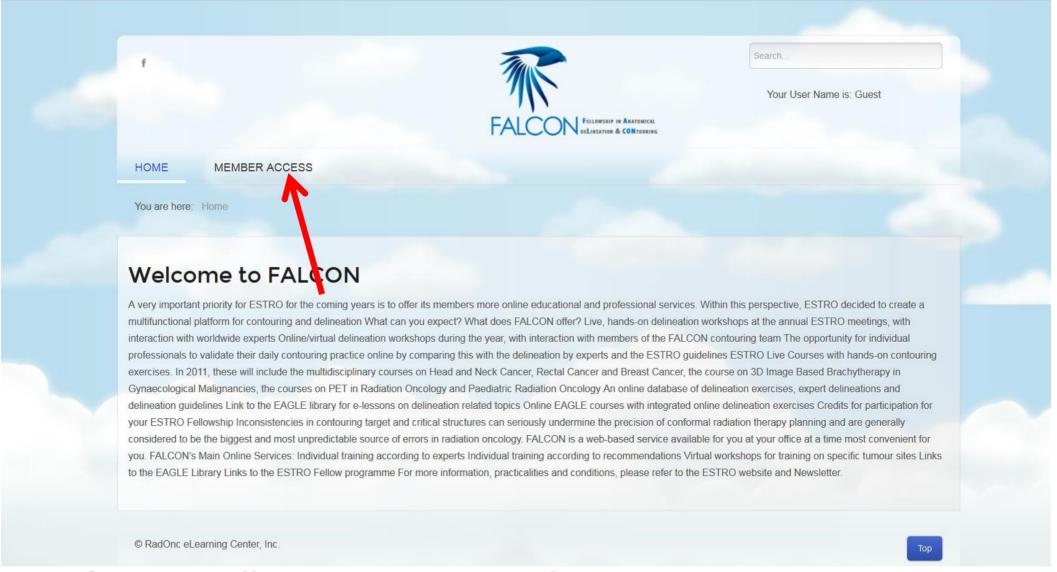


FACON activities

- ESTRO courses
- On site Workshops
- Online Workshops
- Supporting guidelines
- Research projects
 - In pedagogics
 - > In clinical research (Dummy run...)



How to use FALCON?

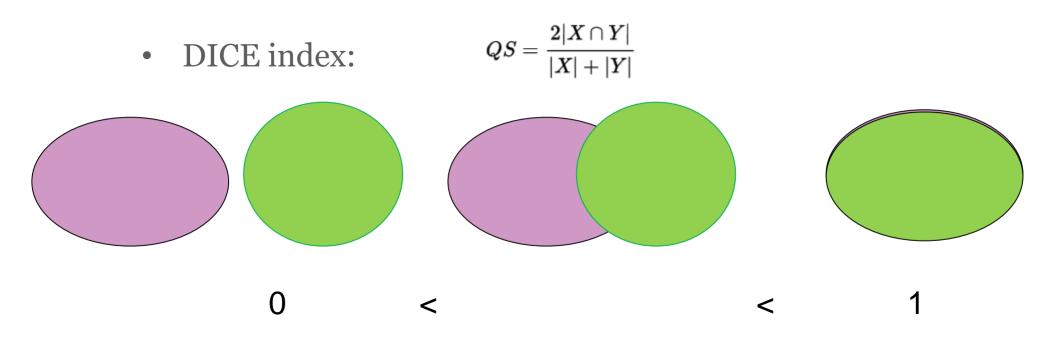


Go to http://estro.educase.com/
Login with your user name and password



Homework results

• 29 participants over 34 did their homework



- Excellent results with DICE index: 0,7 to 0,89!
 - CONGRATULATIONS!!!!



So let's use it again!!!

- OAR contouring in the thoracic region:
 - > Spinal canal
 - > Lung
 - > Heart
 - > Esophagus



Ready to improve your contours?





Image registration

Mirjana Josipovic

Dept. of Oncology, Rigshospitalet Niels Bohr Institute, Uni. of Copenhagen Denmark Peter Remeijer

NKI-AVL Amsterdam The Netherlands

Advanced skills in modern radiotherapy
June 2017



Intended learning outcomes

- Describe basic principles of image registration process
- Identify limitations in image registration process



Image registration

You may also call it

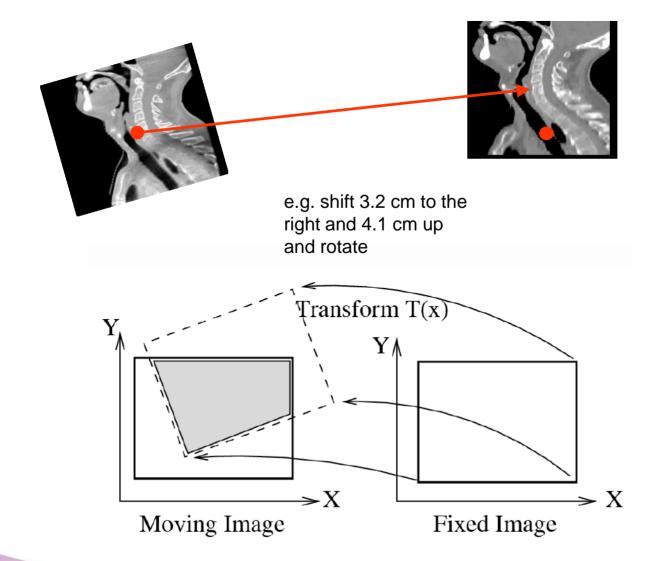
- Image fusion
- Image matching
- Image warping
- = process of aligning two (or more) images





Definition: Image registration

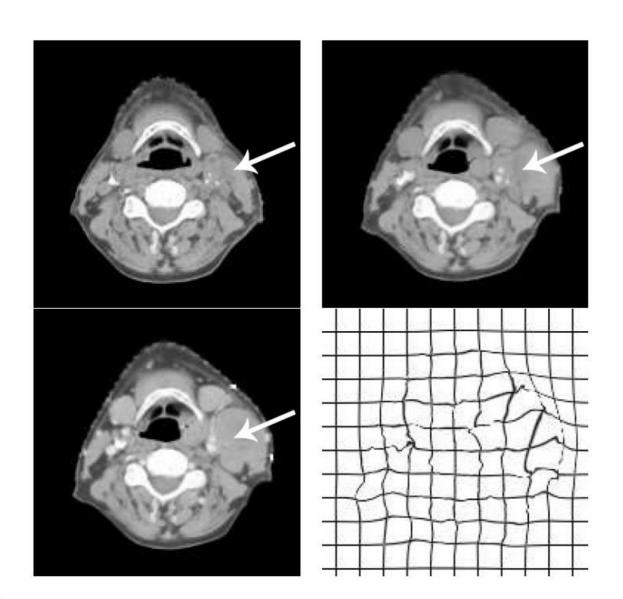
• Determine **rigid transformation** between two scans





Definition: Image registration

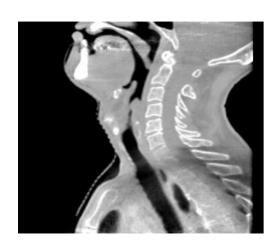
• Determine **deformable transformation** between two scans

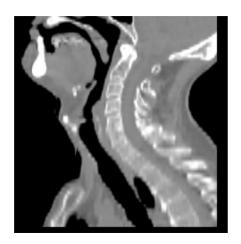


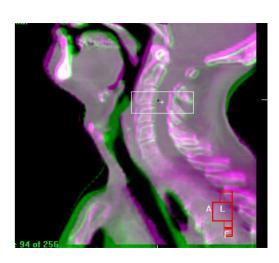


Definition: Image fusion

Combine the information of two images







Viewing and validation of registration result



Image registration in radiotherapy

- In the RT planning process
 - Inclusion of PET/MRI
 - > Pre-chemo CT scans



- Reduction of setup uncertainty
- Detect patient anatomy changes during treatment
 - Daily dose assesment / plan adaptation
- After RT
 - Follow up (tumour response, normal tissue damage)
 - Re-irraditaion





Image registration in radiotherapy

- Algorithms
- Validation
- Challenges





Manual image registration

- Simple 'algorithm'
- Good for gross alignment



- Difficult in 3-D
- Not very precise

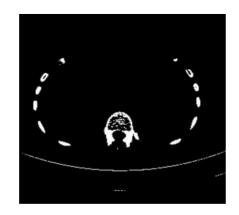




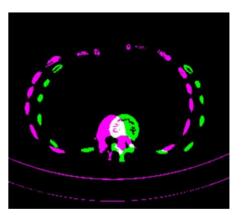
Automatic image registration

- Geometry based
 - Point matching (anatomic landmarks, implanted fiducial markers)
 - Surface matching (skull surface, pelvic bones)
 - Fx Chamfer matching











Automatic image registration

- Feature based
 - Numerical gray scale
- Uses gray values in all pixel values
 - Inside the regions of interest
- Slower than chamfer matching
 - > not really an issue today due to more computing power ©



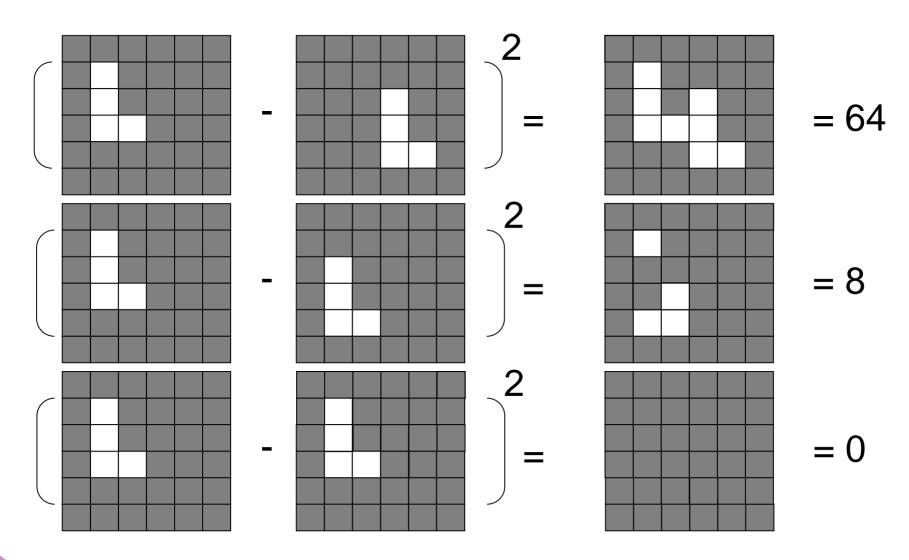


Similarity assessment

- How good is the resulting image registration?
- Similarity metrics / cost functions
 - Root mean square
 - Correlation ratio
 - Mutual information
 - > ...



Grey value registration example



Mutual information cost function



Understand your registration algorithm

• You see:



The computer sees:

$$R = \begin{bmatrix} \cos \theta_z & -\sin \theta_z & 0 \\ \sin \theta_z & \cos \theta_z & 0 \\ 0 & 0 & 1 \end{bmatrix} \begin{bmatrix} \cos \theta_y & 0 & \sin \theta_y \\ 0 & 1 & 0 \\ -\sin \theta_y & 0 & \cos \theta_y \end{bmatrix} \begin{bmatrix} 1 & 0 & 0 \\ 0 & \cos \theta_x & -\sin \theta_x \\ 0 & \sin \theta_x & \cos \theta_x \end{bmatrix}$$

$$= \begin{bmatrix} \cos \theta_y \cos \theta_z & -\cos \theta_x \sin \theta_z + \sin \theta_x \sin \theta_y \cos \theta_z & \sin \theta_x \sin \theta_z + \cos \theta_x \sin \theta_y \cos \theta_z \\ \cos \theta_y \sin \theta_z & \cos \theta_x \cos \theta_z + \sin \theta_x \sin \theta_y \sin \theta_z & -\sin \theta_x \cos \theta_z + \cos \theta_x \sin \theta_y \sin \theta_z \\ -\sin \theta_y & \sin \theta_x \cos \theta_y & \cos \theta_x \cos \theta_y \end{bmatrix}$$

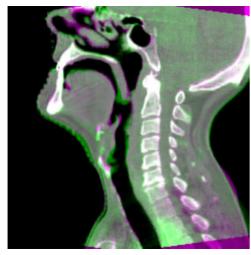
Specify how the algorithm should handle image registration:

- Define region of interest
- Choose the appropriate algorithm

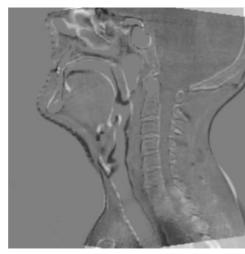
Check the result!



Viewing & validation of the image registration



Overlay



Substract

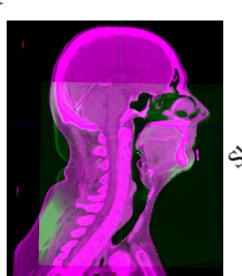


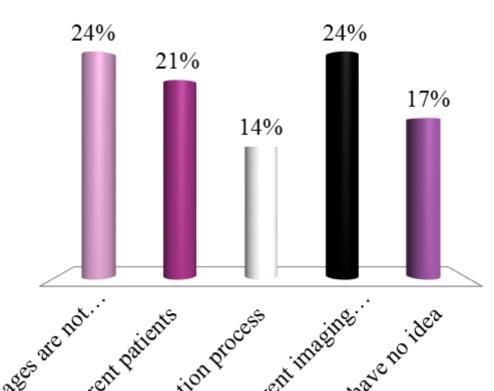
Checker



Why does this overlay look so purple?

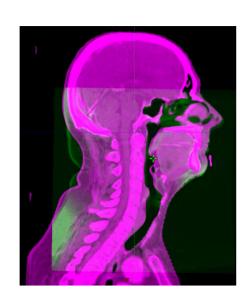
- A. Shown images are not weighted equally
- B. Two different patients
- C. Error in registration process
- D. Two different imaging modalities
- E. I have no idea

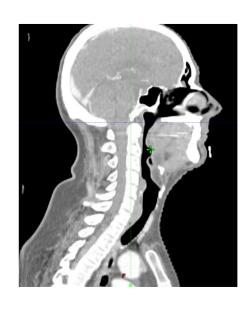


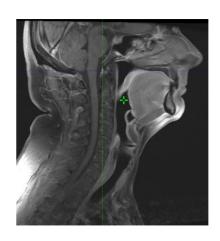




• Viewing & validation of the image registration

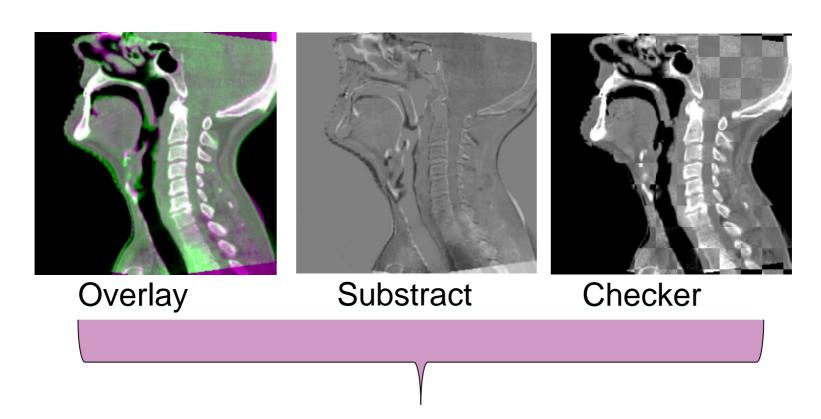








Viewing & validation of the image registration



Same modality



Viewing & validation of the image registration



Split window

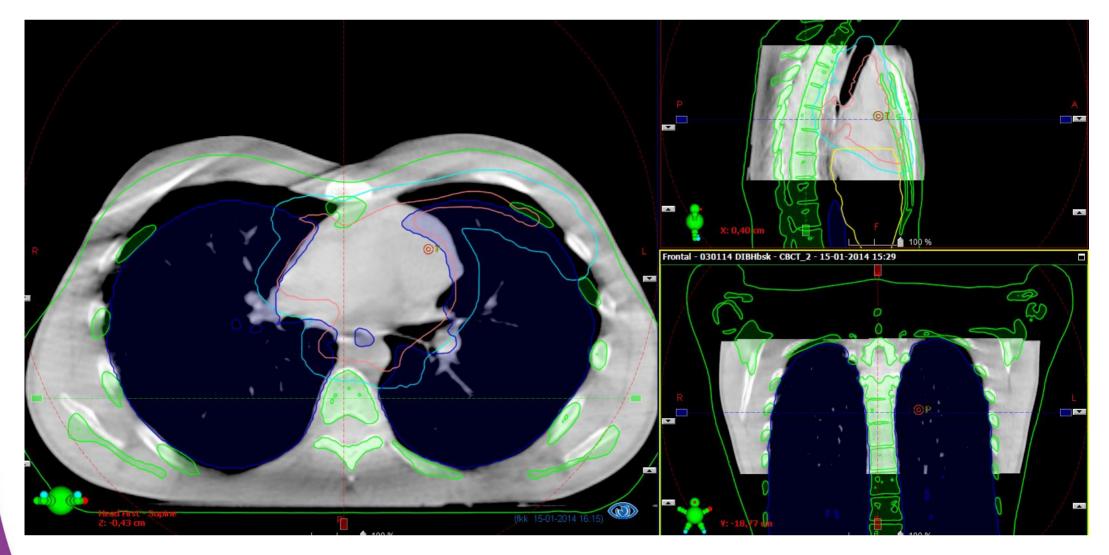


Spy glass

Different modalities



Case: error in automatic image registration

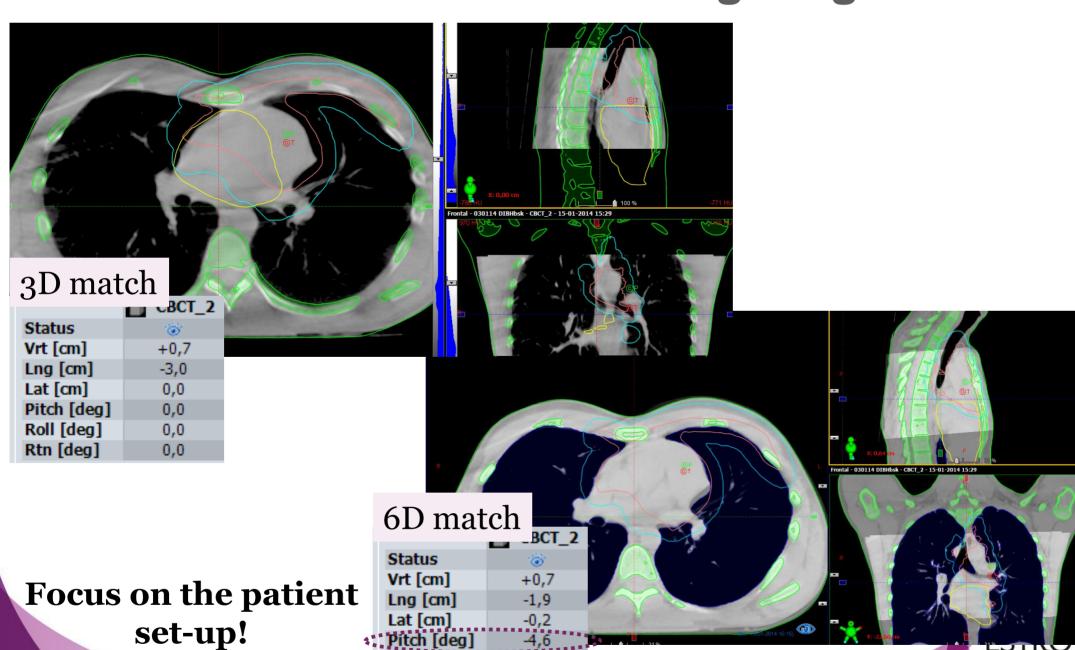


Automatic image registration has to be evaluated!

Focus not only on the primary structures of interest, but on
the whole image!

School

Case: error in automatic image registration



-1,4

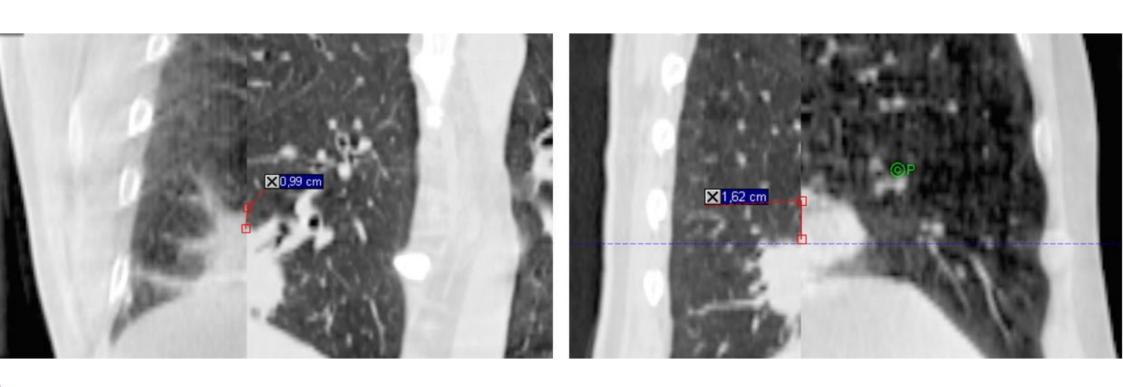
-1,1

School

Roll [deg]

Rtn [deg]

Lung tumour baseline shift



... misalignment of the peripheral tumour after registration on vertebrae



Challenges in image registration





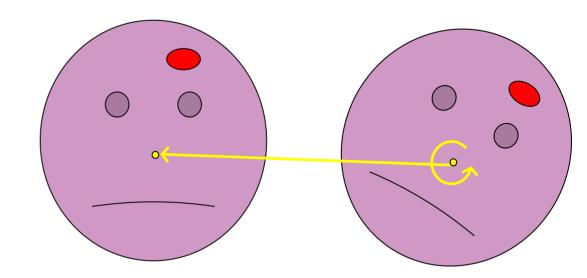
Impact of rotations on image registration

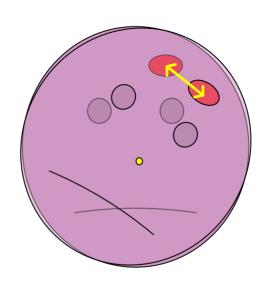
• Registration

- Bony anatomy
- Translations and rotations
- Very accurate



- Only translations
- Potentially large errors



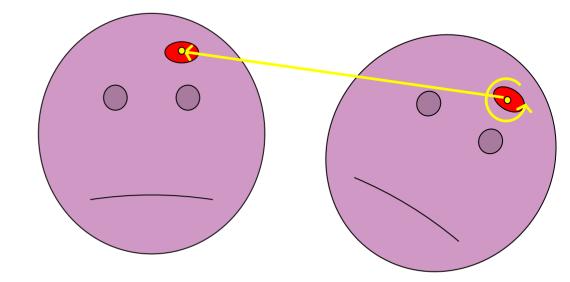




Impact of rotations on image registration

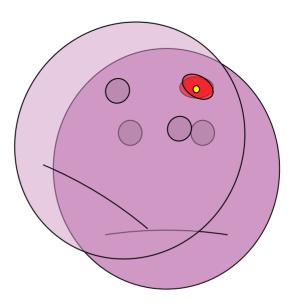
Registration

- Redefine match volume
- Isocenter position



Correction

- Only translations
- Rotational errors are small close to rotation center





Corrections without rotations

Rule of thumb: $\Delta = 0.02 \times \phi \times r \text{ (mm)}$

- 3° rotation
- CTV diameter is 40 mm (r = 20 mm)
- Rotation centre is in CTV $_{\rightarrow}$ Errors to CTV will be smaller than 1 mm $_{r}$ = 20 mm

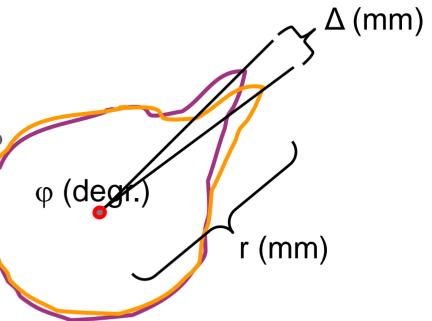


Corrections without rotations

Rule of thumb: $\Delta = 0.02 \times \phi \times r \text{ (mm)}$

Problem for structures far from rotation center

- 3° rotation
- Rotation centre is in CTV
- → 6 mm shift at 10 cm distance!
- → does treatment plan allow this?





Lung stereotactic body radiotherapy

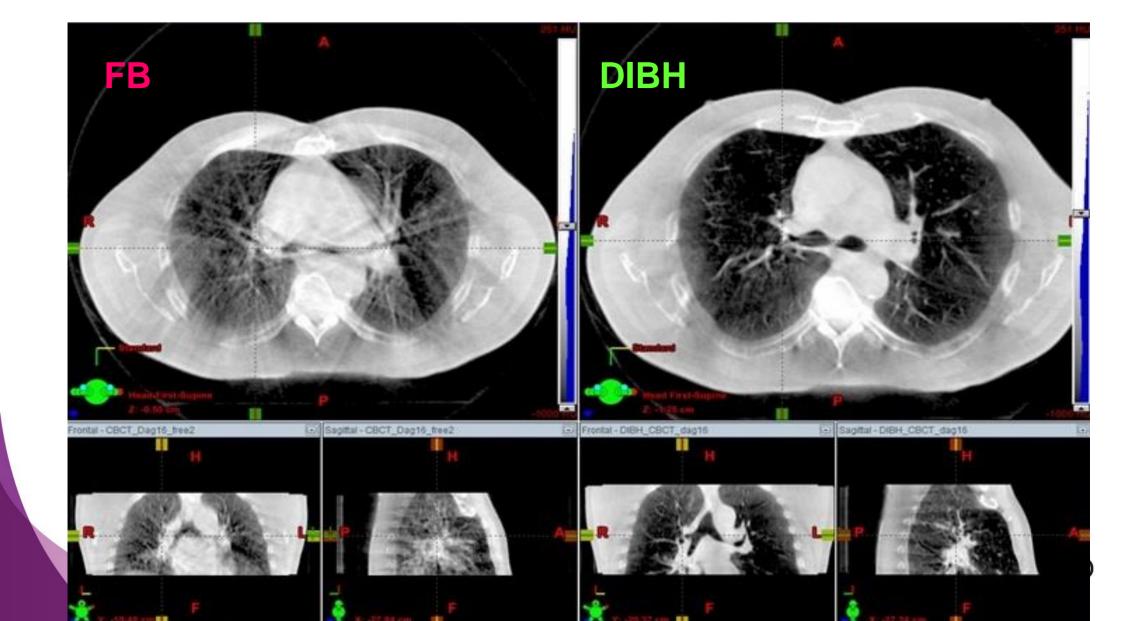
 Residual positional error when only translations were used for image registration

	AP [mm]	CC [mm]	LR [mm]	pitch [°]	roll [°]	rot [°]
M	-0,1	0.1	-0.1	0.31	-0.06	-0.56
Σ	1.3	1.0	0.8	1.20	-0.06 1.23	1.42
σ	1.5	1.0	1.1	0.79	1.33	0.85

AP, anterior-posterior; CC, cranio-caudal; LR, left-right.

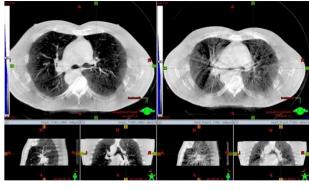


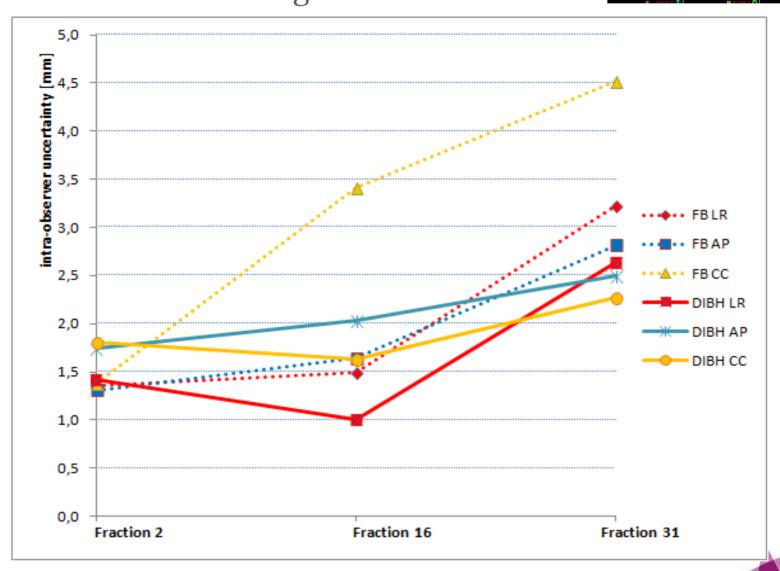
Impact of image quality



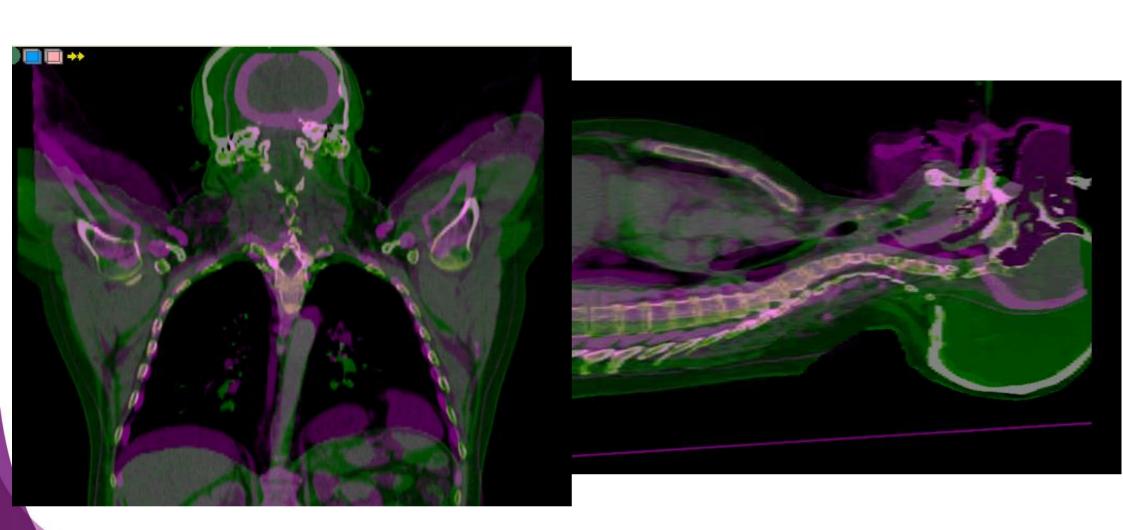
Impact of image quality

Observer uncertainties in CBCT registration





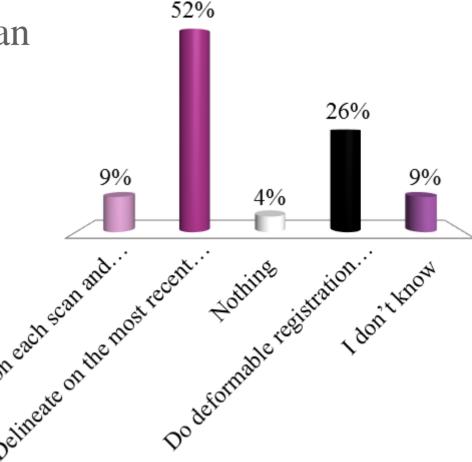
Case: fusion of pre- and post-chemo scan



What would you do (or your radiation oncologist)?

- A. Delineate on each scan and combine contours
- B. Delineate on the most recent scan
- C. Nothing
- D. Do deformable registration before delineation
- E. I don't know





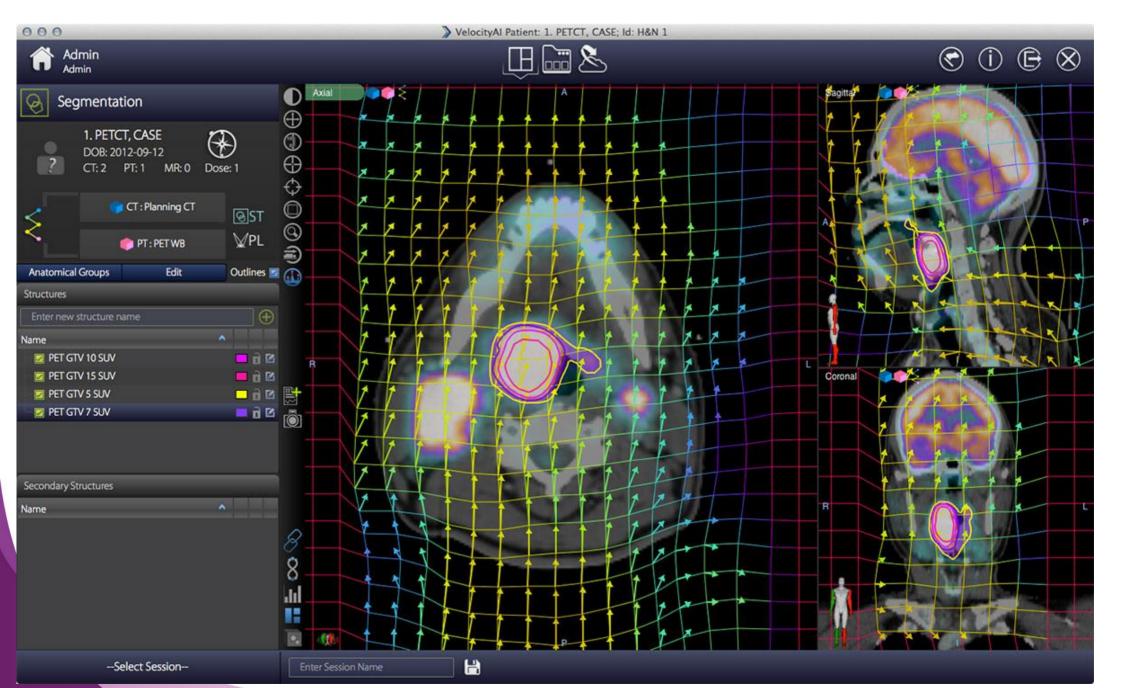


How to handle registration uncertainties?

- Ensure a treatment-like position already at staging
 - > Flat table top
 - Arms up
 - Chest board
 - Motion management
- Good collaboration with the PET / MR department!



Deformable image registration - DIR



Deformable image registration

- How do you know the result is good?
 - ➤ It looks ok ©
- Getting the contours / outlines of organs right
 - Ok for IGRT
- Getting the heterogeniety/tissue cells inside the organs right
 - Necessary for dose accumulation
- Different challenges with different organs
- DIR needs to be evaluated for each clinical problem



Take home messages

- Image registration plays an important role for:
 - routine treatment planning
 - routing treatment delivery
 - Follow up, clinical studies, re-irradiation
- Consider the effect of rotations and anatomical changes
- There is no perfect solution:
 - use best registration algorithm for each problem
 - <u>always</u> include a visual inspection step in the process







Management of respiratory motion in radiation therapy



Mirjana Josipovic

Dept. of Oncology, Rigshospitalet

& Niels Bohr Institute, University of Copenhagen Denmark

Advanced skills in modern radiotherapy
June 2017



Intended learning outcomes

- Differentiate between different motion management strategies in RT
- Interpret the purpose of motion management for different patient groups
- Identify the limitations in motion management



Management of respiratory motion in RT

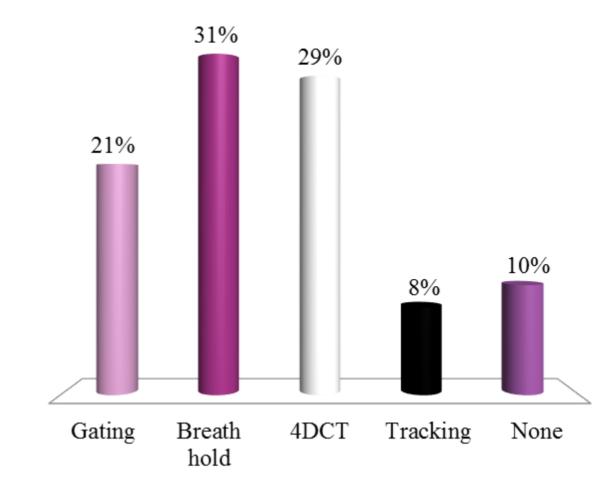
- Respiratory gating technique
- Breath hold methods
- Motion encompassing methods
- Respiration synchronized techniques

AAPM TG 76 definition



Which motion management do you use?

- A. Gating
- B. Breath hold
- C. 4DCT
- D. Tracking
- E. None

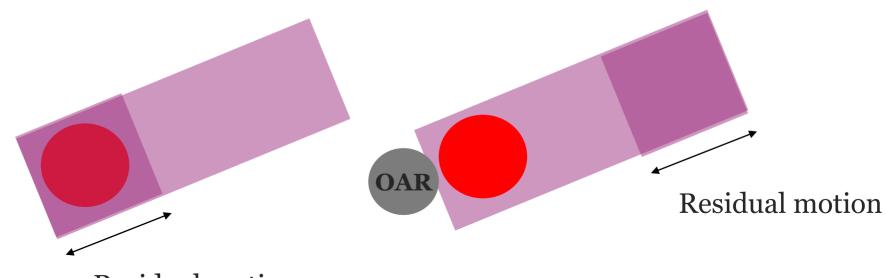


Multiple answers allowed



What is respiratory gating?

 Applying radiation within a particular part of the patient's breathing cycle



Residual motion

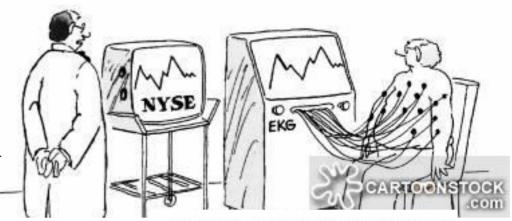
- → Reduce motion during treatment
- → Move target away from OAR



Condition for success with gating

Strong correlation Internal organ motion - External chest motion

- Tumour type and location
- Source of the respiratory signal
- Reproducibility of respiration

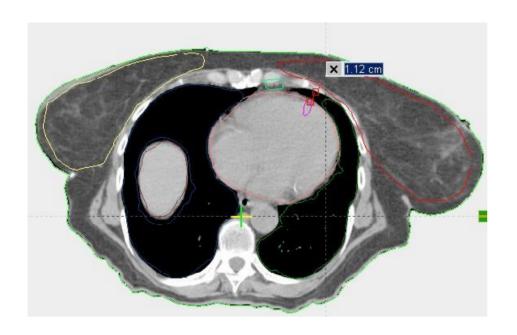


" Amazing . . the patterns are the same! "

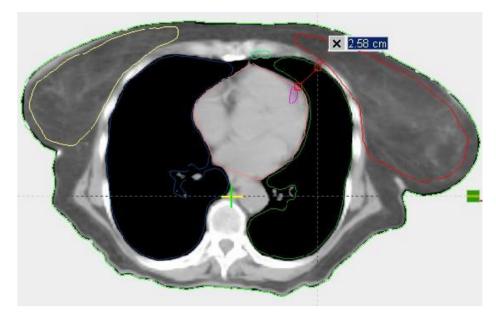


External vs. "internal" motion - breast

Good correlation



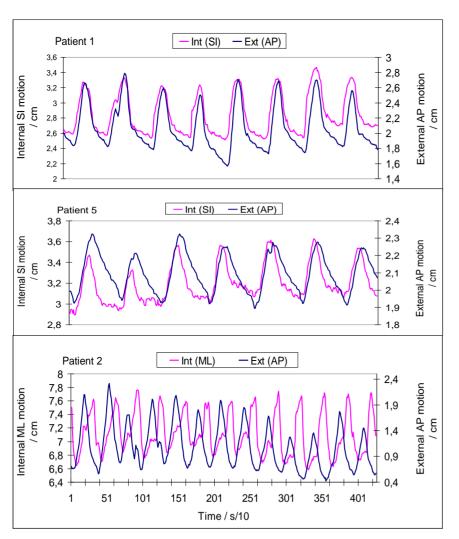
Spontaneous breathing



Enhanced inspiration gating



External vs. internal motion - lung



Correlation can be established

Phase difference

Phase drift

No correlation





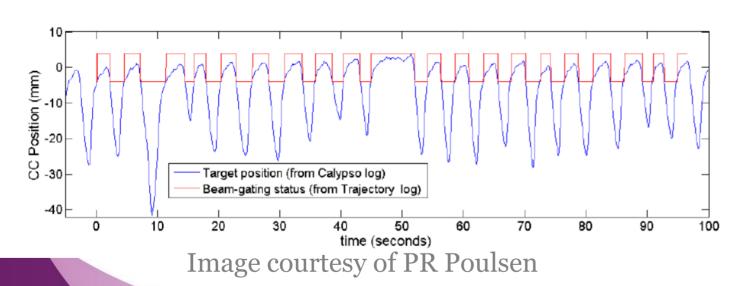
No external vs. internal motion correlation

Simple approach:

Don't do gating

Complicated approach:

Monitor the target position during (gated) treatment







• Free breathing respiratory gating can be applied if there is a good correlation between the respiratory signal and the tumour motion



The choice of breathing technique

Inspiration gating

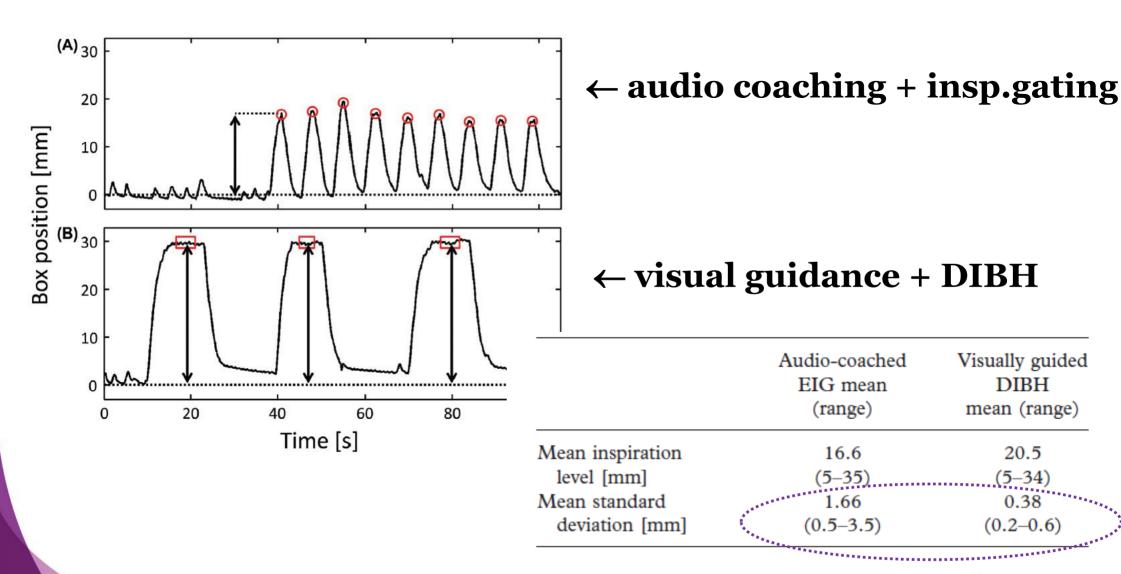


Deep inspiration breath hold (DIBH)





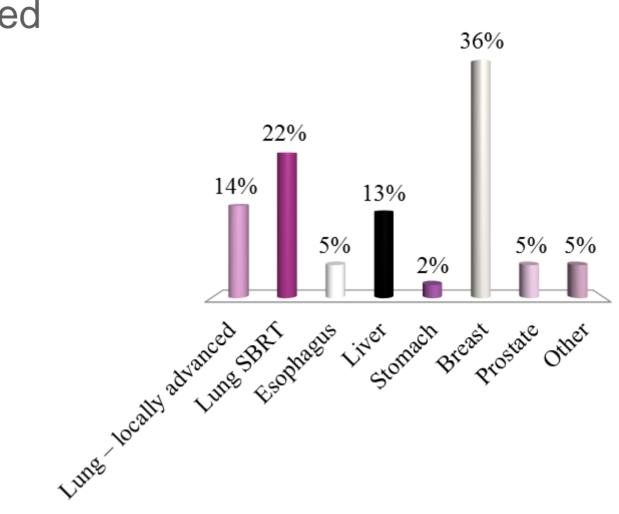
Respiration reprodicibility





In which sites do you use gating / DIBH?

- A. Lung locally advanced
- B. Lung SBRT
- C. Esophagus
- D. Liver
- E. Stomach
- F. Breast
- G. Prostate
- H. Other





How to DIBH?



- Computer-controlled
 - Breathing volume based
 - Optical surface tracking





Breathing volume based DIBH

Spirometry

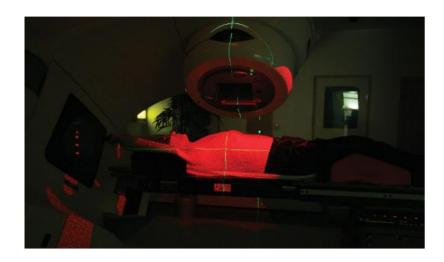


 ABC^{TM}



Optical surface tracking based voluntary DIBH

Surface tracking (Surface Guided RT) Marker tracking



C-RAD SentineI™ Align RT®



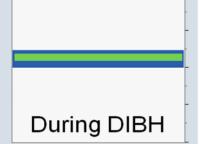


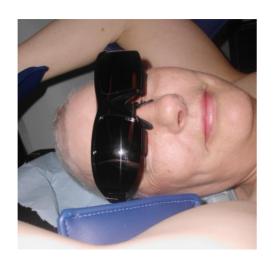


Patient training & QA

- Patient information
- Patient coaching







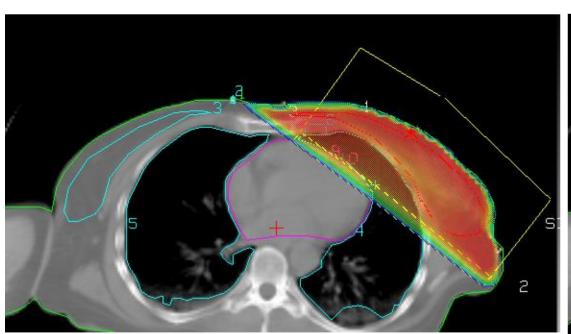


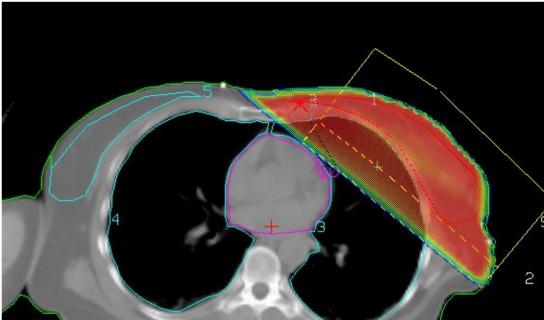


- DIBH level / volume individually adjusted!
- DIBH duration 15-30 s
 - If the patient doesn't comply exclude!



Dosimetric potential of DIBH – breast





Free breathing

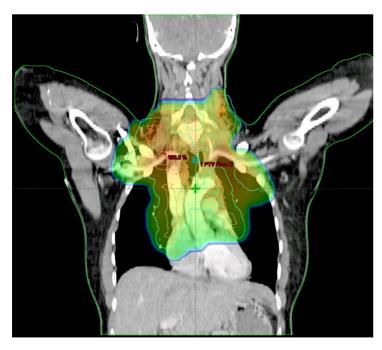
DIBH

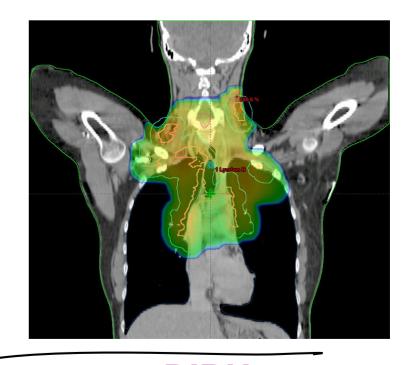
Separation of target / OAR

- Sparing of cardiac structures
- IMN coverage not compromised



Dosimetric potential of DIBH – lymphoma





Free breathing

DIDI

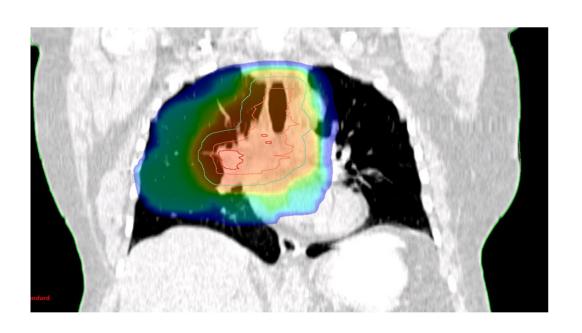


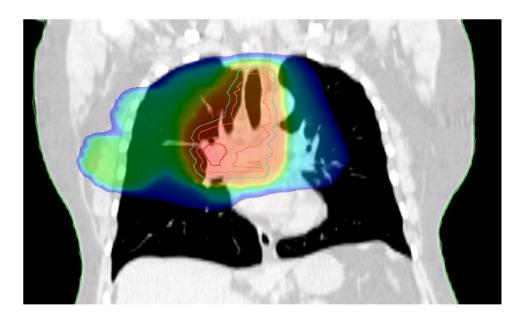
Separation of target & OAR

Images courtesy of Marianne Aznar



Dosimetric potential of DIBH – lung





Free breathing

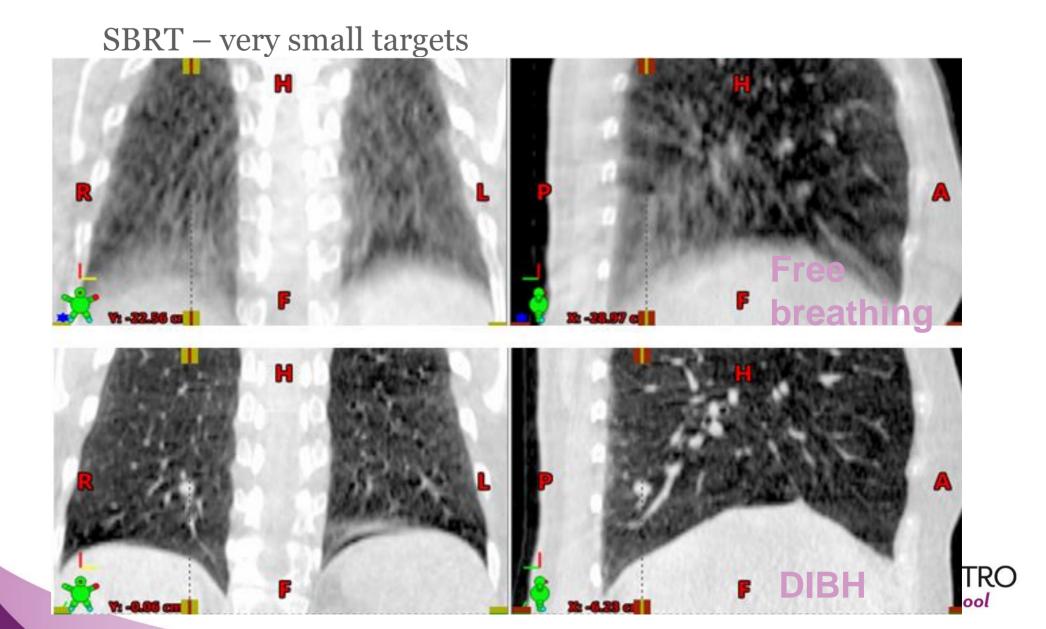


DIBH

- Sparing of heart & lung
- Maintain curative treatment intent
- Tumour motion reduction



Special case: lung SBRT



DIBH gating is more reproducible than inspiration gating

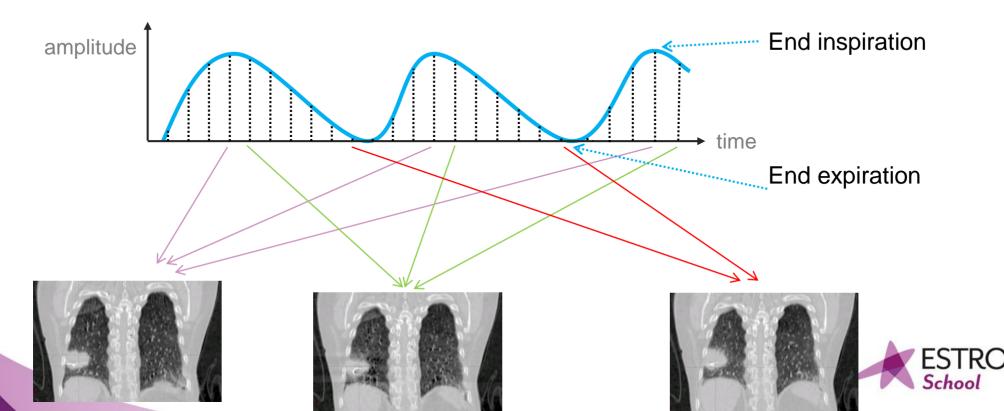
DIBH facilitates anatomical separation of target & OAR

DIBH mitigates target motion



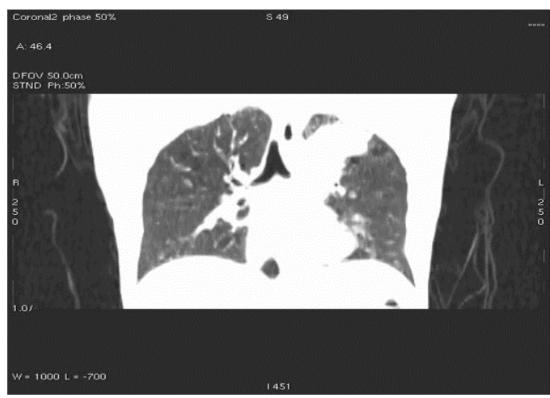
4DCT - a motion encompassing method

- A very slow CT
- Sorting of images acc. to respiration
 - > Resp. phase
 - Resp. amplitude



4DCT – a motion encompassing method





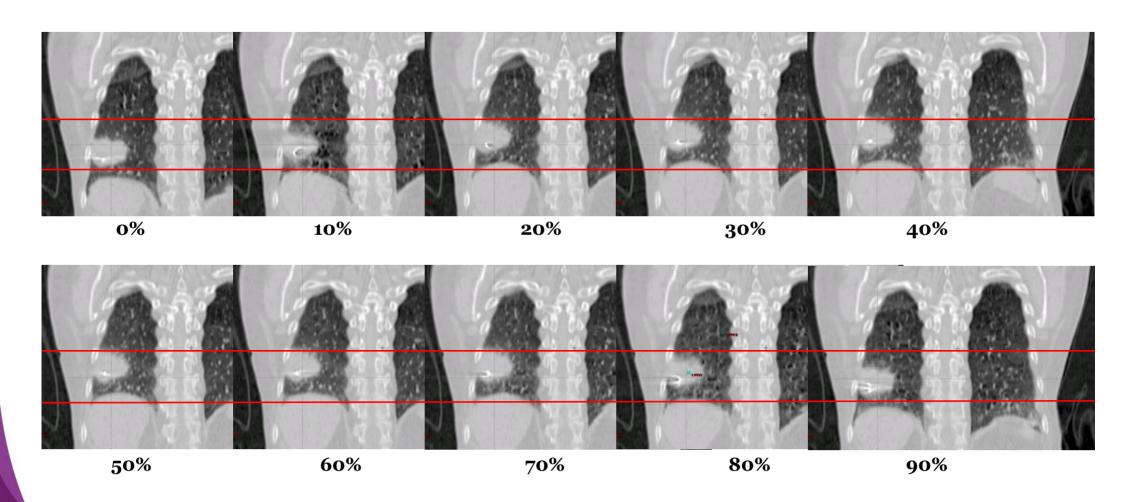


4DCT facilitates

- Tumour motion evaluation
- Delineation of ITV or
- Selection of midventilation phase
- Correlation of tumour position respiratory phase
- 4DCT is only a snapshot!



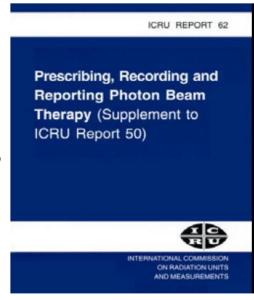
4DCT = 10 3D CTs from 10 respiratory phases

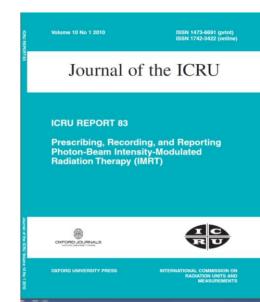




ITV = internal target volume

- ITV = margin for tumor motion added to CTV
- *ICRU 62: "ITV = CTV + margin for uncertainties in size, shape & position of CTV within the patient"*
- iGTV = sum of GTVs in al phases of 4DCT
- ICRU 83: "resulting PTVs were too big"



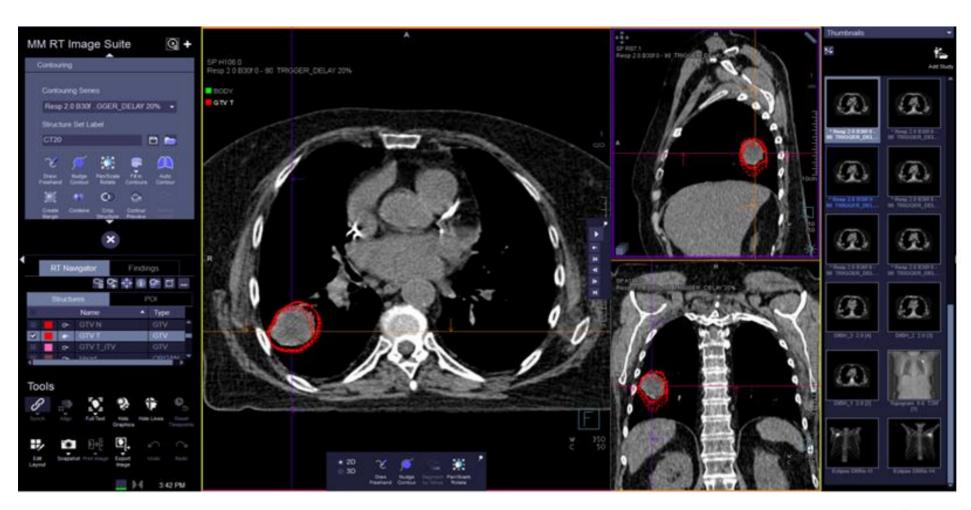








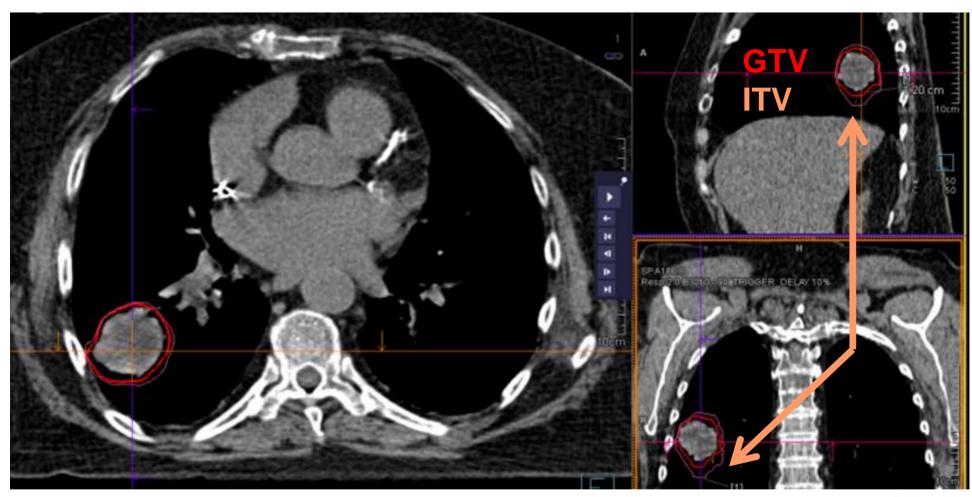














4DCT: Midventilation



Int. J. Radiation Oncology Biol. Phys., Vol. 65, No. 5, pp. 1560–1571, 2006

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0360-3016/06/\$-see front matter

doi:10.1016/j.ijrobp.2006.04.031

PHYSICS CONTRIBUTION

MID-VENTILATION CT SCAN CONSTRUCTION FROM FOUR-DIMENSIONAL RESPIRATION-CORRELATED CT SCANS FOR RADIOTHERAPY PLANNING OF LUNG CANCER PATIENTS

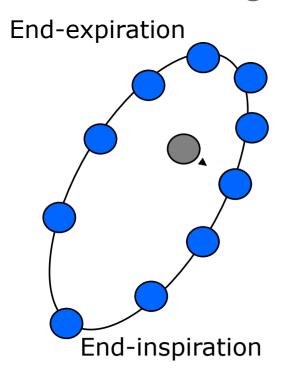
Jochem W. H. Wolthaus, M.Sc., Christoph Schneider, Ph.D., Jan-Jakob Sonke, Ph.D., Marcel van Herk, Ph.D., José S. A. Belderbos, M.D., Maddalena M. G. Rossi, D.C.R.(R), R.T.T., Joos V. Lebesque, M.D., Ph.D., and Eugène M. F. Damen, Ph.D.

Department of Radiation Oncology, The Netherlands Cancer Institute, Antoni van Leeuwenhoek Hospital, Amsterdam, The Netherlands

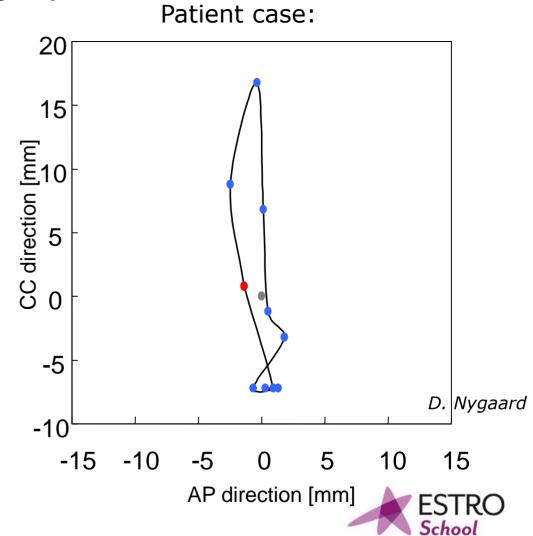
Purpose: Four-dimensional (4D) respiration-correlated imaging techniques can be used to obtain (respiration) artifact-free computed tomography (CT) images of the thorax. Current radiotherapy planning systems, however, do not accommodate 4D-CT data. The purpose of this study was to develop a simple, new concept to incorporate patient-specific motion information, using 4D-CT scans, in the radiotherapy planning process of lung cancer patients to enable smaller error margins.

4DCT: Midventilation

= time weighted average position of tumour

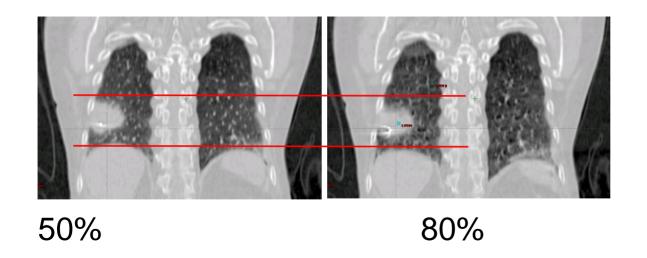


- Tumour position in phases 1-10
- Time weighted average tumor position
- Midventilation

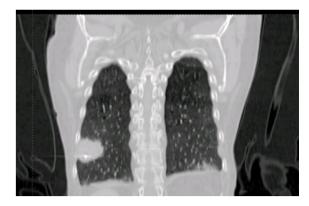


Midventilation

- choice of the correct phase



• Comparisson of tumour size & shape with the breath hold scan



Breath Hold scan



Midventilation vs. midposition

MIDVENTILATION = 1 phase of the 4DCT

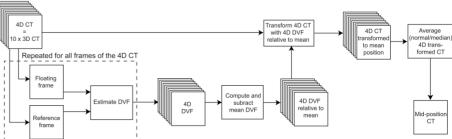


MIDPOSITION =

Deformable registration

Deforming phases to time-weighted midposition

Averaging (median)

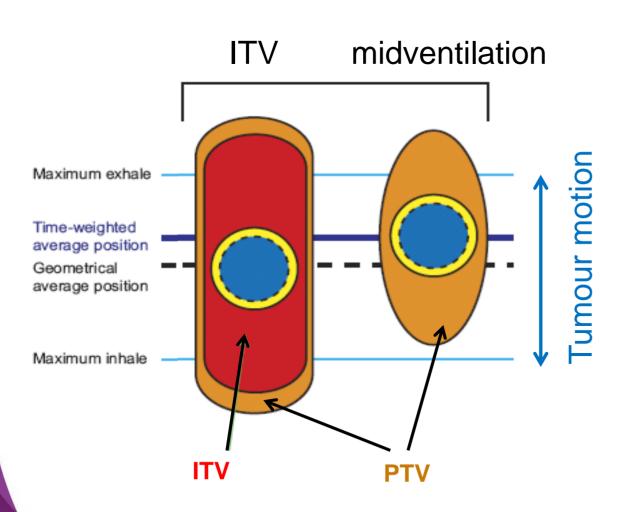


Wolthaus 2008

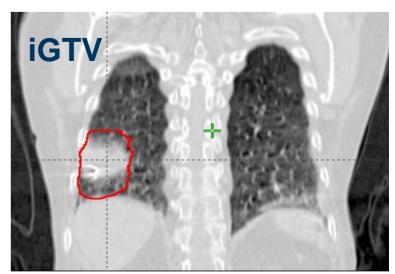


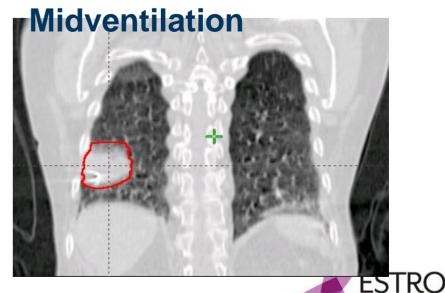
Images courtesy of Marcel van Herk

ITV or midventilation – impact on PTV



adapted from J Wolthaus IJROBP 2008





ITV-like approach

- Larger GTV
- Smaller GTV-PTV margin
- Larger PTV
- Beneficial if hysteresis in tumour motion

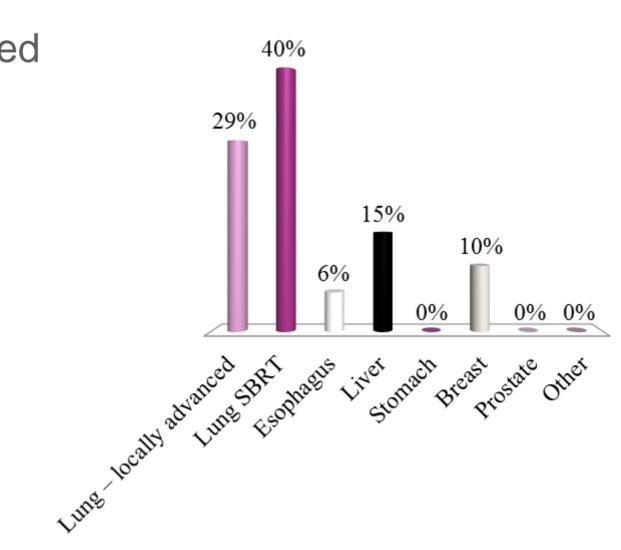
Midventilation

- Smaller GTV
- Larger GTV-PTV margin
- Smaller PTV
- Problem if hysteresis in tumour motion



In which sites do you use 4DCT?

- A. Lung locally advanced
- B. Lung SBRT
- C. Esophagus
- D. Liver
- E. Stomach
- F. Breast
- G. Prostate
- H. Other





Respiration synchronised techniques

Rationale of motion tracking...

Letting the beam move with the target

How

- By using surrogate for tumor motion:
 - external or internal
- Prediction algorithms

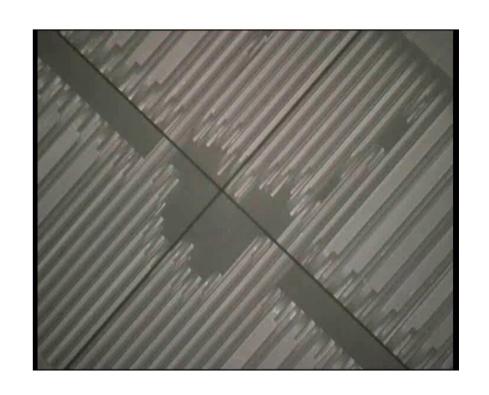




Respiration synchronised techniques

Tracking on linac

• MLC shape adjusted to compensate for target motion in real-time

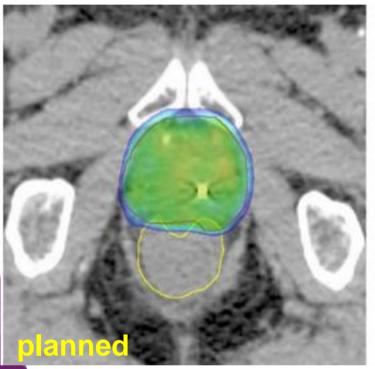


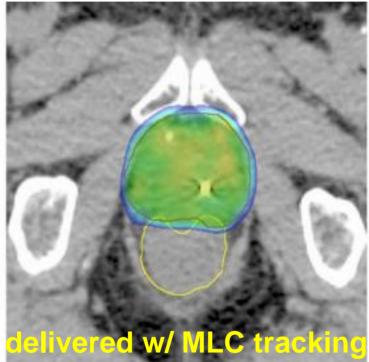
You need to KNOW the target motion!

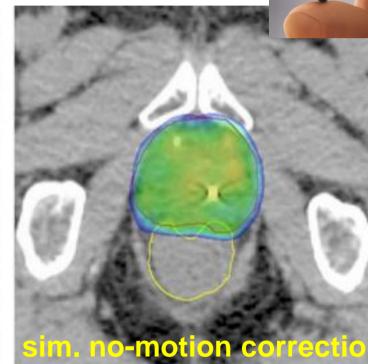


Motion synchronised techniques

First patient treated with electromagnetic transponder MLC tracking











Take home messages

- Different motion management strategies
 - Gating
 - Breath hold
 - Tracking
 - 4D imaging
- Good correlation between respiration surrogate & target motion

Dosimetric benefit!

Patient training improves reproducibility





Treatment Planning



Liz Forde, MSc (RTT)
Assistant Professor
Discipline of Radiation Therapy
Trinity College Dublin





Trinity College Dublin

Coláiste na Tríonóide, Baile Átha Cliath The University of Dublin

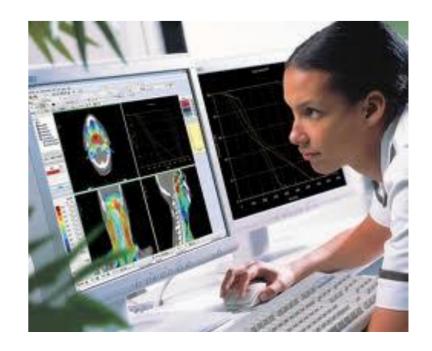
Learning Outcomes

- Discuss the role of the RTT in the treatment planning process
- Discuss key concepts of ICRU 50, 62 and 83
- Identify key features of inverse planning techniques
 - > IMRT
 - > VMAT
- Identify evidence for the use of inverse planning
- Describe the inverse planning process for IMRT and VMAT
- Describe the importance of target and organ definition and it's impact on the inverse planning process
- Review the benefits of inverse planning to "non standard" sites



RTT Lead Planning

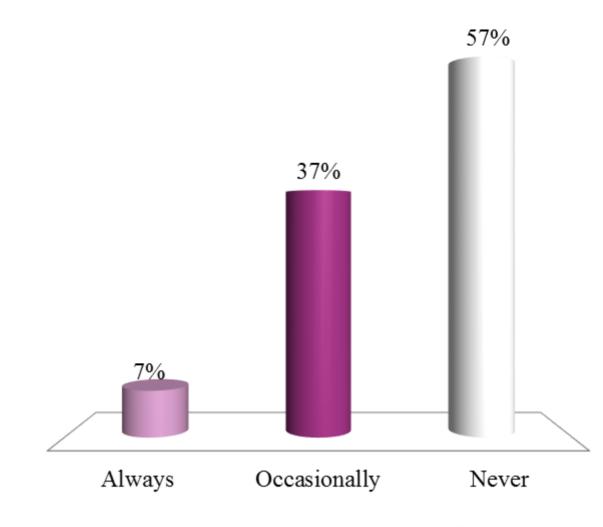
- Scope of practice may vary significantly
- Often seen as a "Specialist role"
 - Rotations may be limited
- Regardless of level of involvement in planning, a basic understanding of key principles increase your "clinical intelligence"





In my work, I am involved in treatment planning:

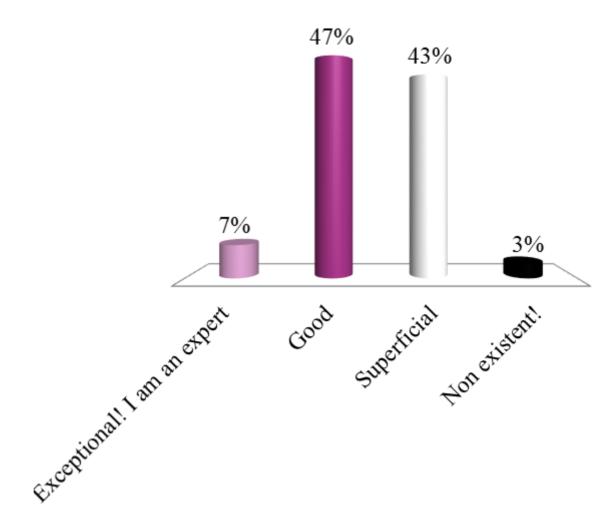
- A. Always
- B. Occasionally
- C. Never





My knowledge and understanding of treatment planning and theoretical concepts is:

- A. Exceptional! I am an expert
- B. Good
- C. Superficial
- D. Non existent!

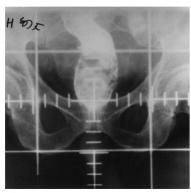


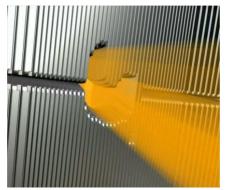


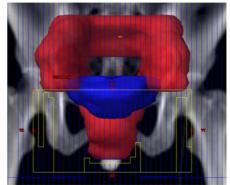
Planning: Where are we now?

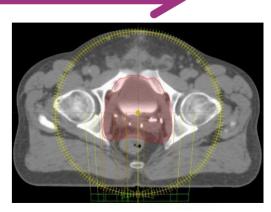
- Technology boom
- From 2D to 3D
- From 3D to 4D, ART and tumour tracking
- From block shielding to conformal shielding
- · From conformal shielding to dynamic shielding
- Inverse planning allows for greater control











Inverse Planning: Is the "Evidence" There?

- Understanding the Literature and the Evidence
- Caution!
 - > Small patient numbers
 - Retrospective in nature
 - Important to recognise fundamental differences in planning techniques between centres
 - Target dose and coverage stipulated
 - o ICRU Pt or Volumetric
 - OAR constraints (protocol or department specific)
 - Beam energy
 - Number of fields/arcs
 - Planning system used
 - Sliding window vs. step and shoot IMRT





"it is important that clear well defined unambiguous, and universally accepted concepts and terminology are used to ensure a common understanding" (ICRU 62)



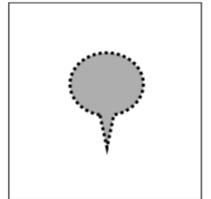


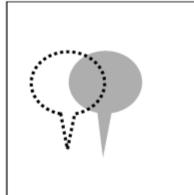
ICRU 50

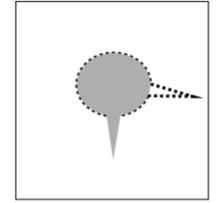
- GTV
- CTV
- PTV
- Irradiated Volume
- Treated Volume
- OAR
- ICRU reference point
- Dose heterogeneity
- (>95%, <107%)

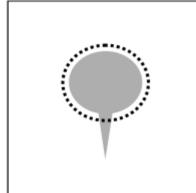
ICRU 62

- Reference points
- Coordinate Systems
- PRV
- ITV
- CI









Feuvret et al., 2006



ICRU 50

- GTV
- CTV
- PTV
- Irradiated Volume
- Treated Volume
- OAR
- ICRU reference point
- Dose heterogeneity
- (>95%, <107%)

ICRU 62

- Reference points
- Coordinate Systems
- PRV
- ITV
- CI

ICRU 83

- Detailed labelling of structures
- Volumetric prescription
- Median dose (D50%)
- Near min (D98%)
- Near max (D2%)
- CI (again)
- HI



The Need for Standardised Reporting

- Green Journal Editorial, 2013 (Yartsev, Muren and Thwaites)
- Planning papers are interesting to everyone (RO, MP and RTTs)
- Pick up practical tips and share outcome data BUT...

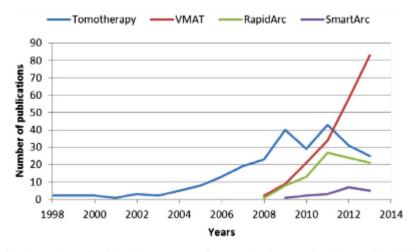


Fig. 1. Number of publications per year for rotational IMRT planning studies. (The data for 2013 is limited to 11 months.)

A third group of readers includes treatment planners who need solid information about the details of the planning procedure applicable to the current case. Unfortunately, there is a variety of definitions and a confusion in terminology that makes it difficult to compare publications of plans performed by different groups. For example, we have found nine different definitions used to describe conformity of the prescribed dose to the target, and seventeen (!) for the homogeneity of dose distribution within the target. The included DVHs should be reproduced in high-quality, allowing for exact numerical values to be derived. It is also essen-



Adherence to ICRU 83 Reporting

- 48 IMRT or VMAT papers published from 2010 2015 were analysed (*A Mohan and E Forde, 2017 Publication currently under review*)
- 22.9% reported PTV D2%
- 18.8% reported PTV D98%
- 8.3% reported PTV D50%

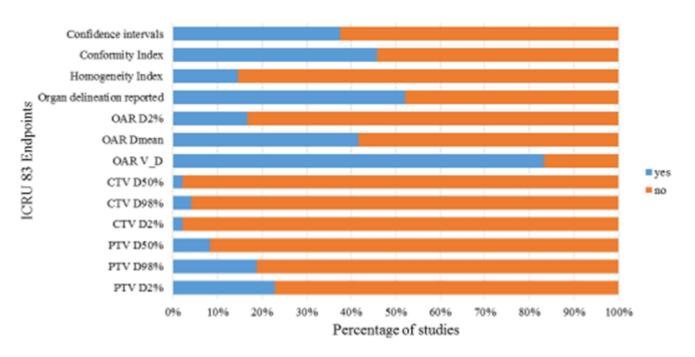


Figure 1 shows the proportion of studies which followed the ICRU 83 Report recommendations for modulated plans.



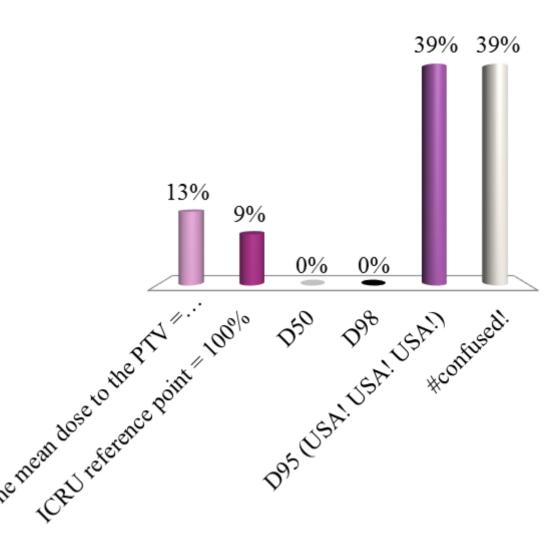
Fine, But What is Happening in Clinical Practice?

- Survey of 10 Academic Institutions in The US (Das et al., **2017**)
- "Nearly 95% of patient treatments deviated from the ICRU-83 recommended D50 prescription dose delivery."
- The majority of institutions appear to be prescribing to D95



In my department we prescribe our IMRT/VMAT prostate plans to:

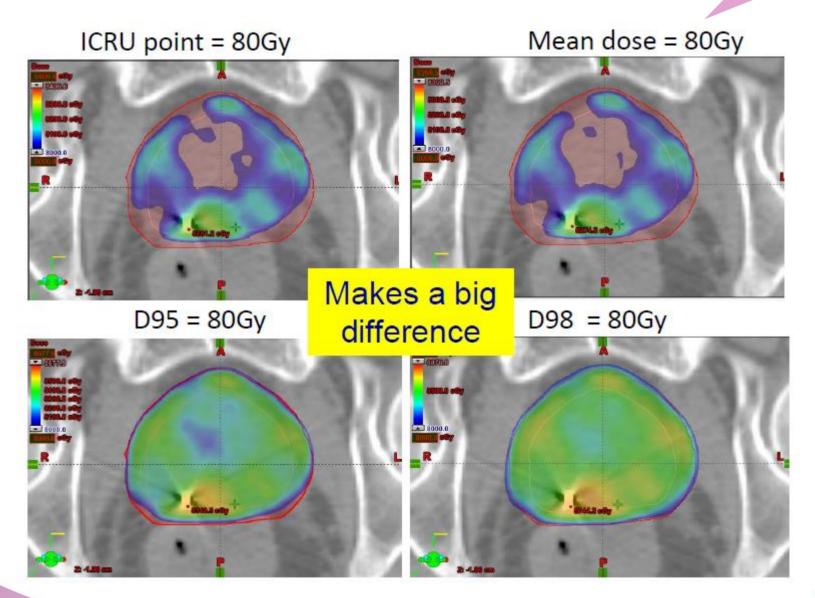
- A. The mean dose to the PTV = 100%
- B. ICRU reference point = 100%
- C. D50
- D. D98
- E. D95 (USA! USA! USA!)
- F. #confused!





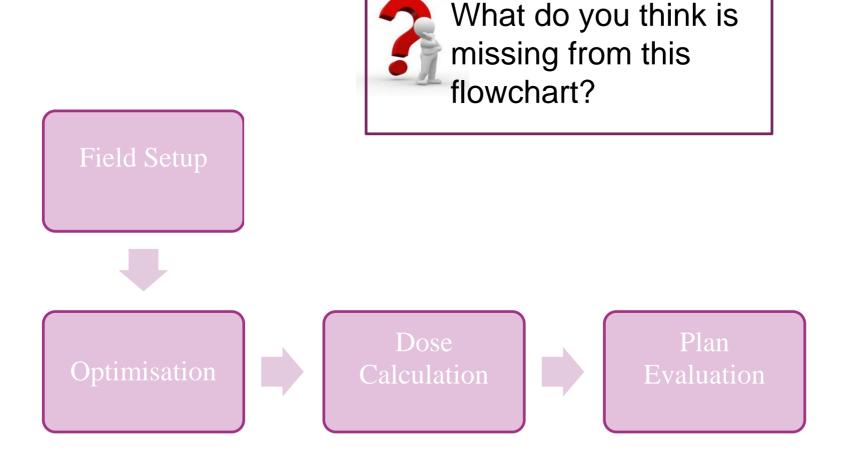
Plan Normalisation

What happened to ICRU 83?



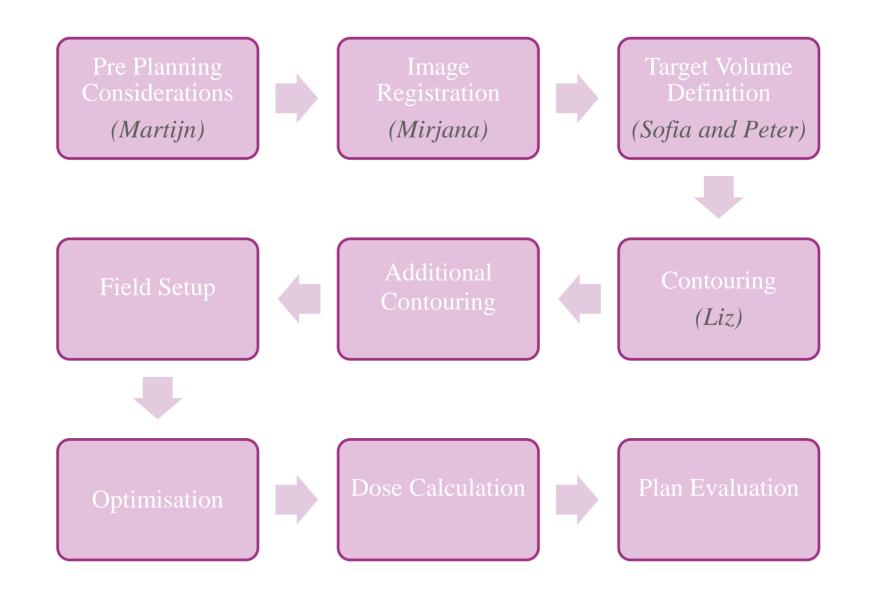


The Planning Process







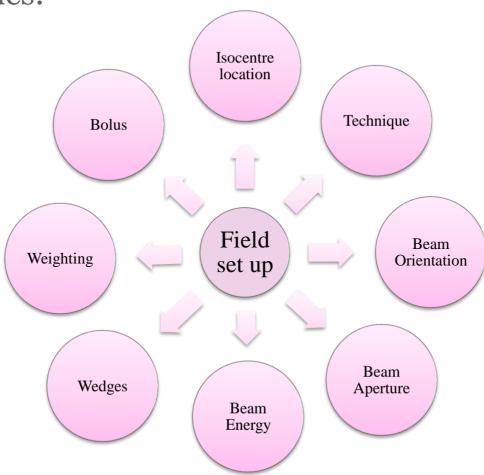




Key Concepts of 3DCRT

• Field set up... "Finally we get to put some beams on!"

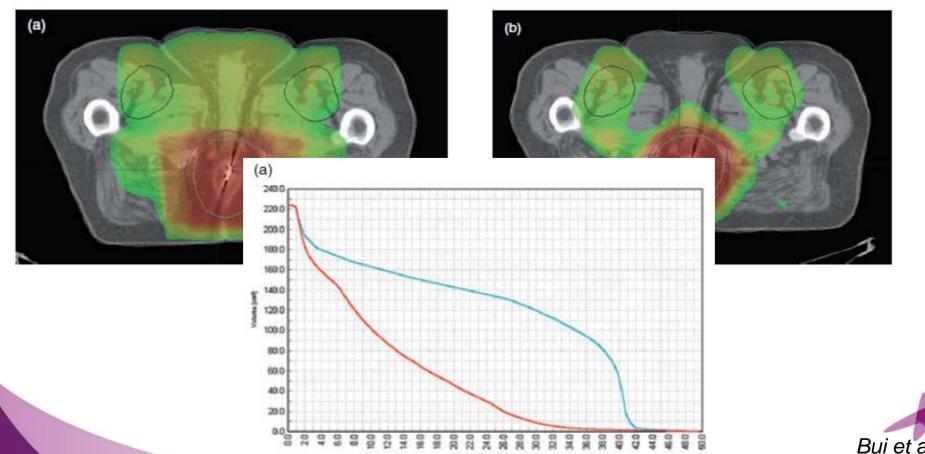
• User defines:





Planning Techniques Explored...

- With 3D targets now being delineated, 3DCRT techniques have become more complex
- "Genital sparing" technique



Key Concepts of 3DCRT

- But...
- How many fields are we up to now?
- Enter IMRT...

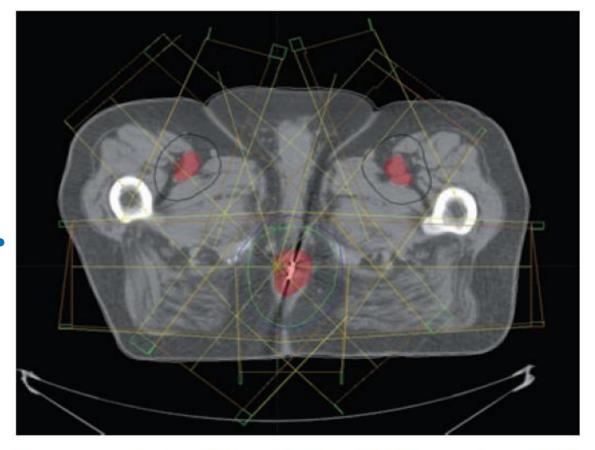


Fig. 2. Example of three-dimensional conformal radiotherapy technique field arrangement. Gross tumour anus and inguinal, red; planning target volume (PTV) anus, green; PTV inguinal, black; PTV pelvis, dark blue.

Bui et al., 2009



- The multiple-static-field MLC technique
 - Step and Shoot
- The dynamic MLC technique
 - Sliding Window
- Intensity modulated arc therapy
 - > IMAT
- Intensity modulated proton therapy
 - > IMPT
- "IMRT requires expertise and careful target design to avoid reduction in local control by marginal miss" (NCCN 2013)



- IMRT is the delivery of radiation to the patient via fields that have a non-uniform radiation distribution across a field.
- Progression from geometric to **fluence** shaping of a field

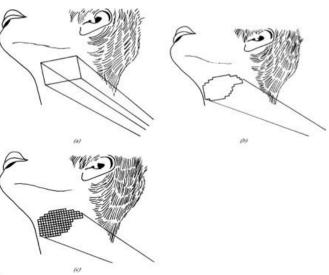
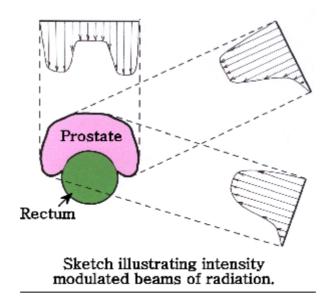
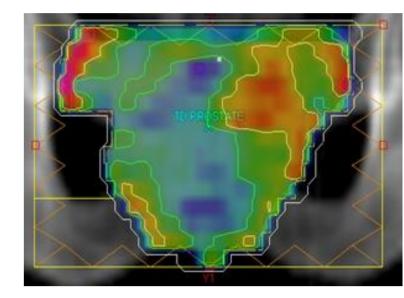


Image taken from: S Webb (2003) The physical basis of IMRT and inverse planning British Journal of Radiology 76: 678-689







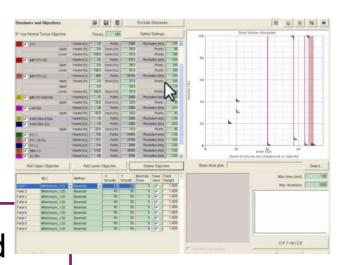
- This fluence is modulated
- The intensity of the fluence changes across the beam
- This changing intensity is based on the required dose to be delivered across a field
- This modulated fluence will determine the dMLC leaf motion

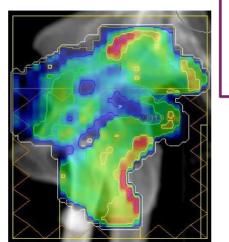


Limitations of IMRT...

- 1. Multiple PTVs
- 2. Complex PTVs (close to skin edge)
- 3. Multiple OARs with multiple DVCs

Sophisticated optimisation parameters





Complex fluence patterns

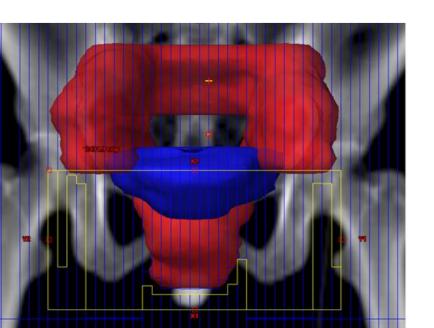


High MUs

• Limitatio SMRT...

Large PTVs

ncreased number of planning fields

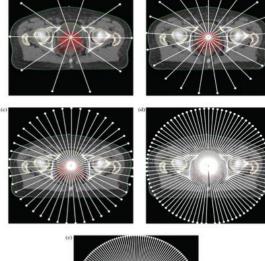


Due to restrictions I motion for SW IMRT, even more treatment fields (for some Varian machines)

NB: this image demonstrates the concept of split carriages

Key Concepts of VMAT

- Simultaneously changing 3 main features
 - MLC leaf motion
 - Gantry speed
 - Variably dose rate
- Inverse planning based on Progressive Resolution Optimisation Algorithm (PRO)
- PRO 3
 - > 4 multi resolution levels
 - All 178 control points are included in each level
 - Internal logic
 - > Intermediate dose calculation



Clinical Applications of VMAT

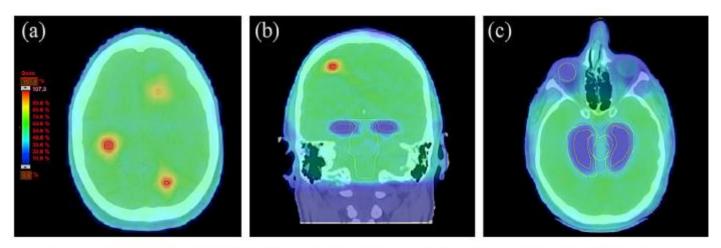
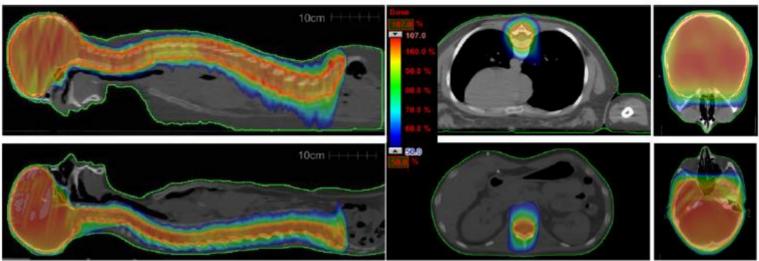


Fig. 1. Examples of isodose distributions for whole brain radiotherapy with hippocampal avoidance and simultaneous integrated boost for three brain metastases using volumetric modulated arc therapy. The whole brain clinical target volume was prescribed to 32.25 Gy in 15 fractions. Three metastases were prescribed 70.8 Gy in 15 fractions. (a) Axial image with three metastases. (b) Coronal image with one metastasis and the hippocampi. (c) Axial image with the hippocampi and eyes.

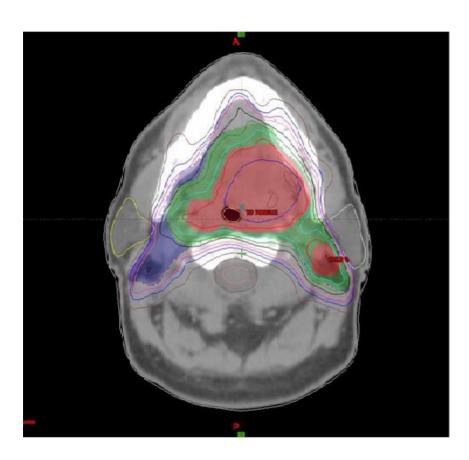
Hsu et al., 2010



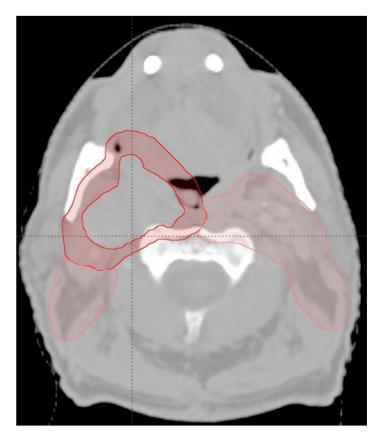
Fogliata et al., 2011



The Benefits of Inverse Planning



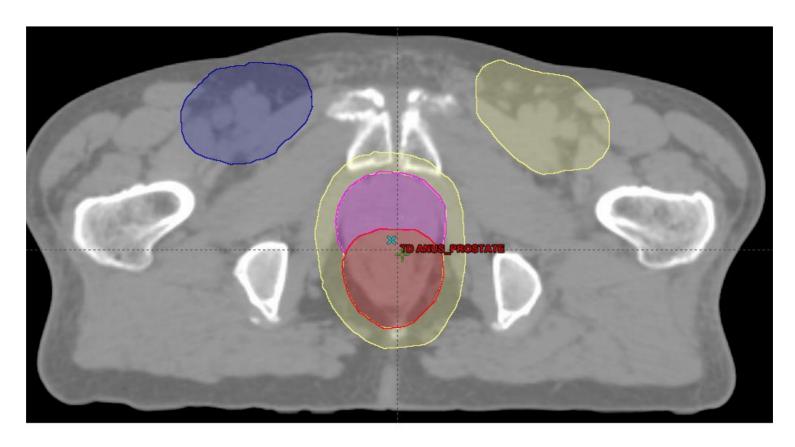
Complex concave volumes



Increased control over distribution
Boosting targets within targets



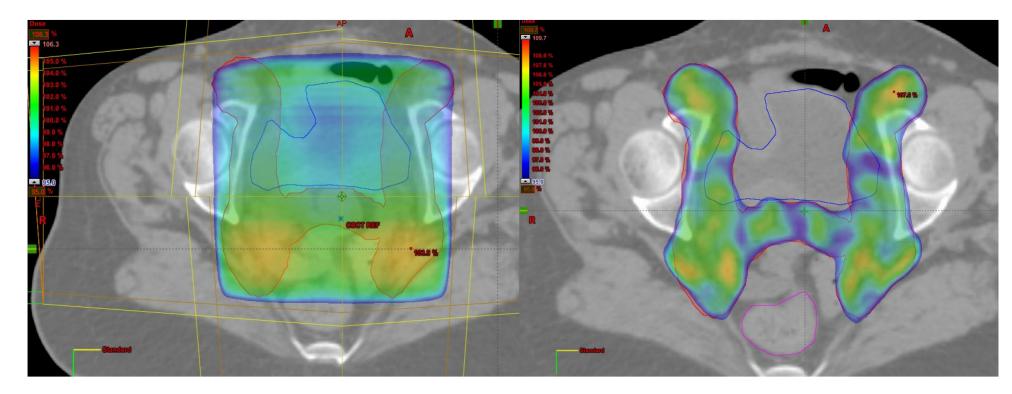
The Benefits of Inverse Planning



Multiple targets
Simultaneous integrated boost



The Benefits of Inverse Planning



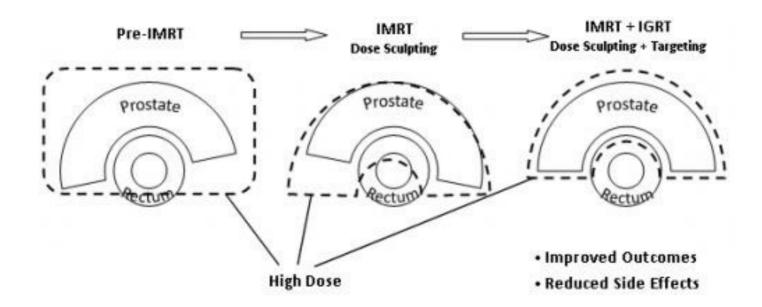
Sharp dose fall off Improved OAR sparing

Need robust IGRT!



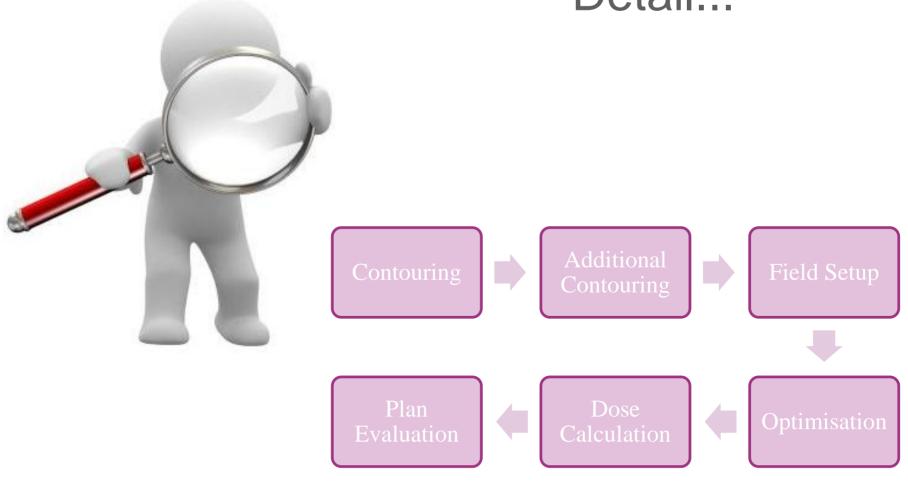
The Benefits of *IG*-IMRT

• Jose will cover this in more detail this afternoon!



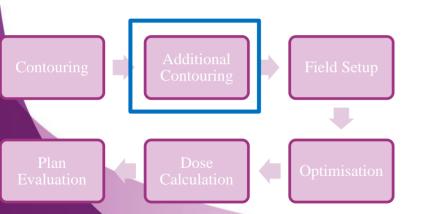


Let's Look at the Inverse Planning Process in Closer Detail...



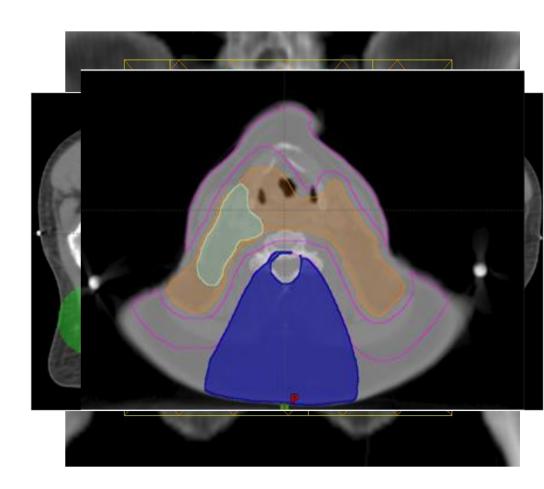


- Virtual contours used only in optimisation but <u>not</u> plan evaluation
- Ease the optimisation process/algorithm
- How and when you use them will depend on the case and also on your experience as a planner
 - ➤ Also what point of the optimisation process you are at for VMAT





- Increase control over dose distribution
 - Dose escalate within a PTV
 - Dose fall off across a structure
 - Dose directly surrounding PTV
 - Dose dumping in healthy tissue





- Improve coverage of whole or partial target
 - ➤ We can't manually adjust the MLC
 - > Inverse planning is volume based planning
 - Can be "cold" on superior or inferior slices
 - Can be "cold" where there is a competition between structures
 - > "IMRT PTV"

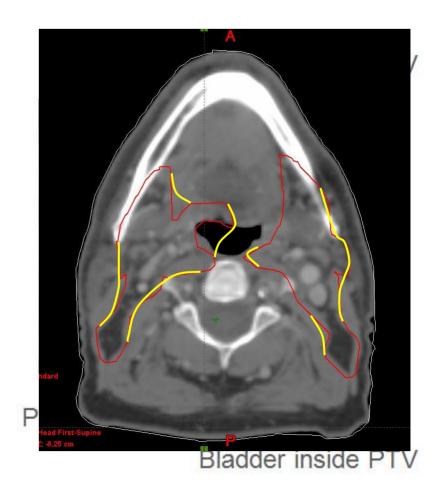


"True" PTV
Used for plan evaluation

"IMRT" PTV
Used for optimisation



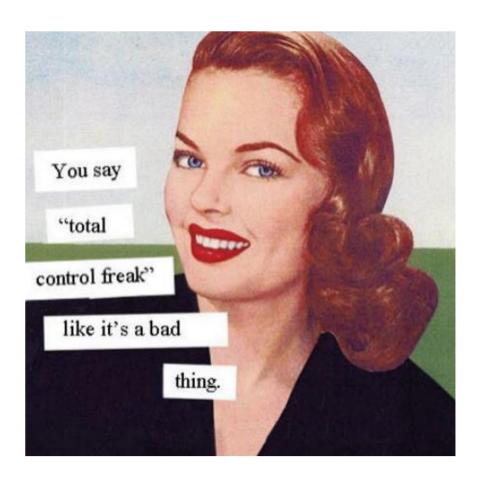
- Lessen the competition between structures
 - OAR and target
- Smoother contours and gradients between slices of target structure





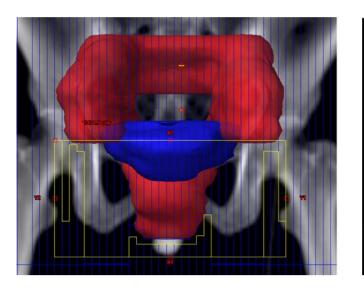
Field Setup

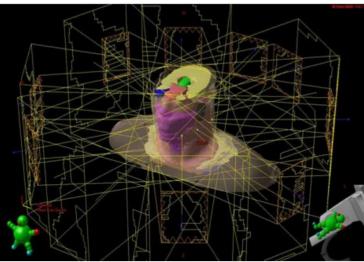
- Isocentre placement
- Beam arrangement
- Field size
- Collimator angle
- Dose rate

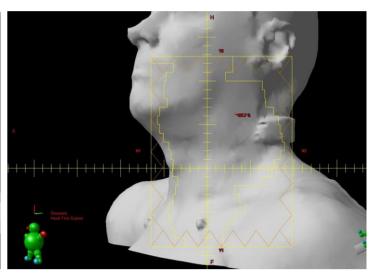




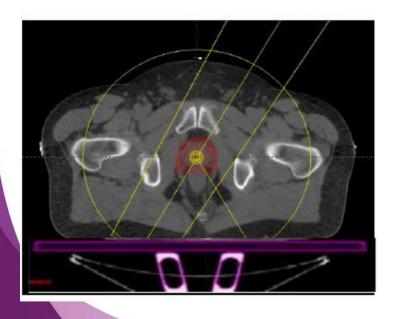
IMRT

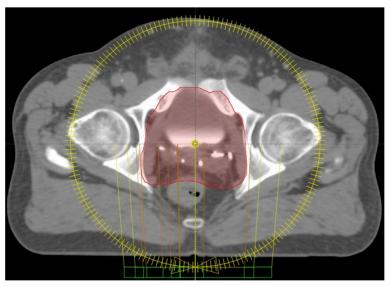






VMAT



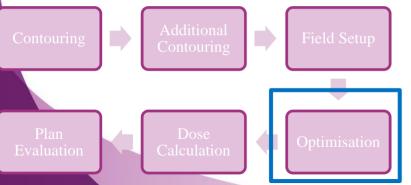


Shoulders: Angle gantry to avoid Angle couch to avoid Fix jaw to avoid (sup or ant/post)



Inverse Planning Optimisation

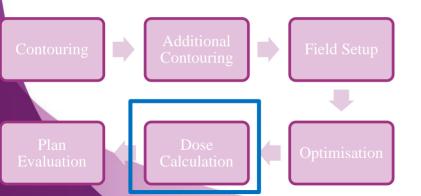
- Planner decides on required dose coverage with dose constraints for surrounding structures
 - Cost function algorithm
- Upper and lower dose limits are to be nominated
 - > Target structures have both
- Planning systems allow for dose constraints to be specified
 - Either as a dose max, mean dose or as a %volume to receive a specified dose
 - Can have either a single point, a series of points or a line





IMRT Dose Calculation

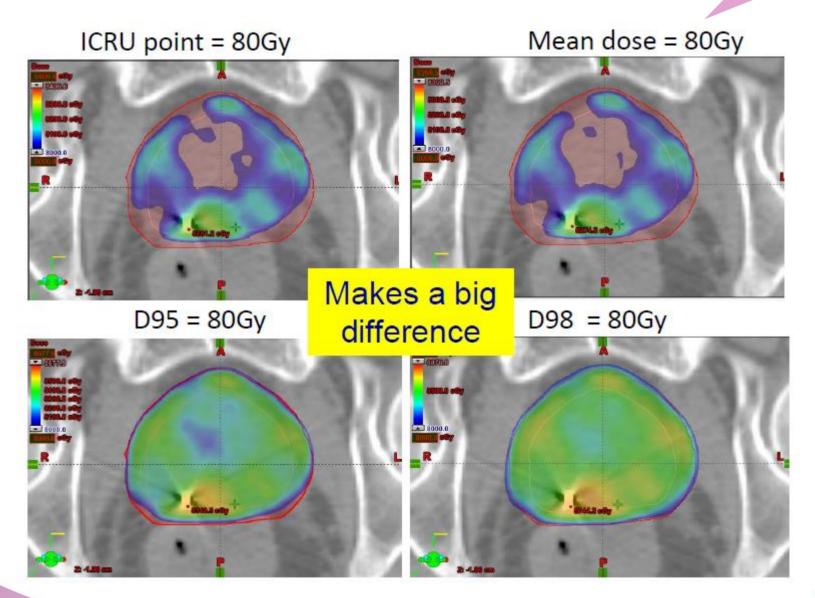
- The fluence maps are generated at the time of optimisation
- The leaf motion is then calculated to enable the delivery of this
- The 3D dose calculation is then carried out generating a dose distribution
- Note the subtle changes:
 - Fluence now reflected the deliverable values
 - ➤ The DVH is now based on AAA as opposed to PBC





Plan Normalisation

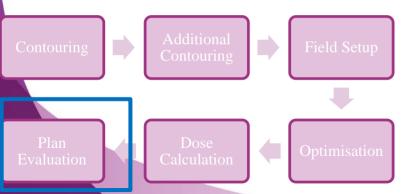
What happened to ICRU 83?





 This is a crucial component of the planning process and should not be rushed or undervalued

- Target Coverage
- Target Conformity
- Target Homogeneity
- OAR doses
- Integral Dose
- Field arrangement used
- Fluence maps or segments for IMRT
- Monitor Units
- Treatment time





- Select appropriate tools
 - Modern TPS are
 developed to make our
 life easier but are only as
 good as the user who is
 interpreting the
 information
- Qualitative
 - Visual inspection is vital
 - Clinical judgement
- Quantitative
 - > ICRU 56, 62, 83
 - > DVH
 - Conformity and homogeneity indices

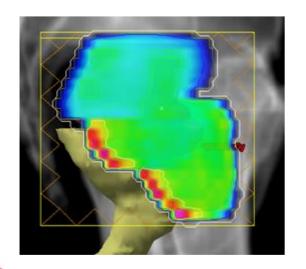


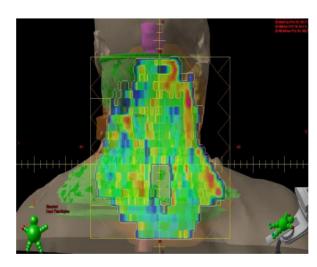
Revise ICRU!

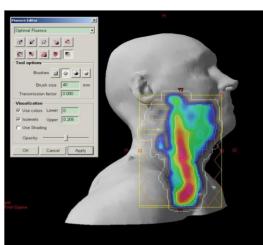
You must know and use the correct terminology You must know the main recommendations



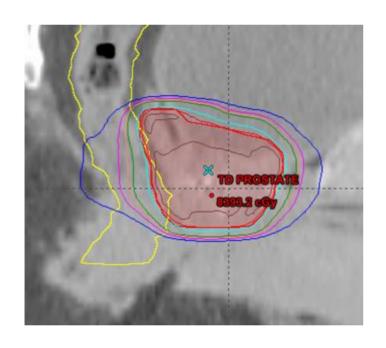
- RTTs care about fluence maps too!
- What is level of modulation
- Is this necessary
- What impact does this have on the dose distribution
- What impact does this have on treatment delivery

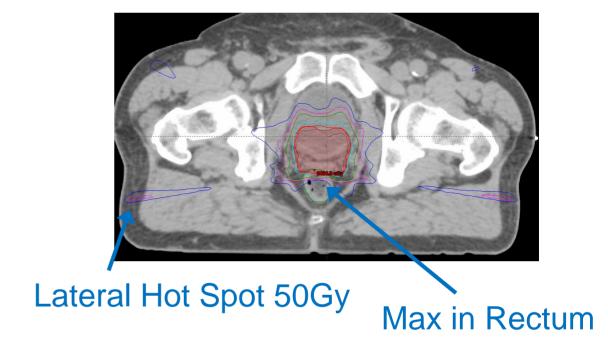


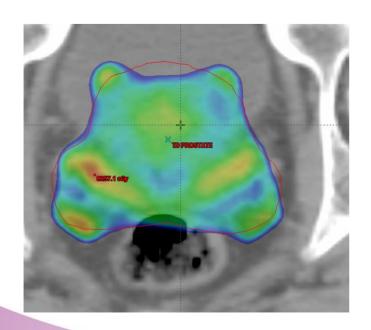


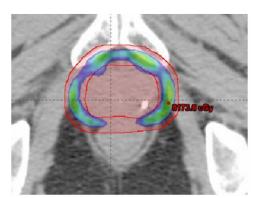


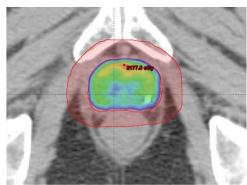














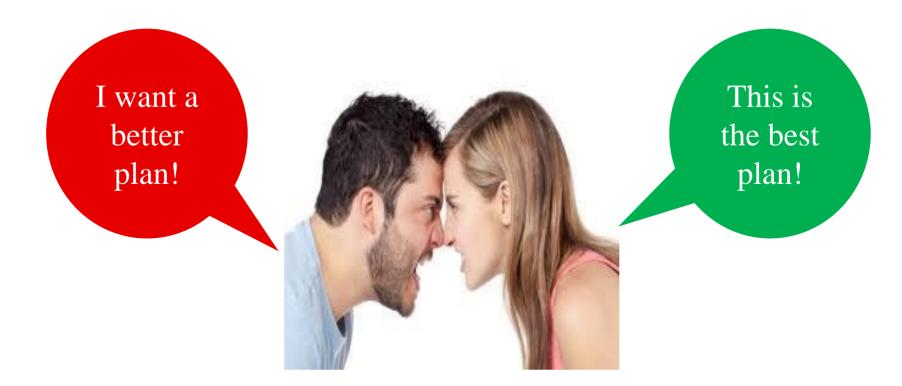
How To Improve a Bad Plan

- Beam Angles
 - Number and position
 - > Bare in mind length of treatment
- Plan normalisation
 - Heat up or cool down the whole plan
 - Quick, does not require re calc
- Reoptimise
 - Think about what you are trying to achieve
 - Relax constraints if possible
 - > Try to keep it simple



Just Remember...

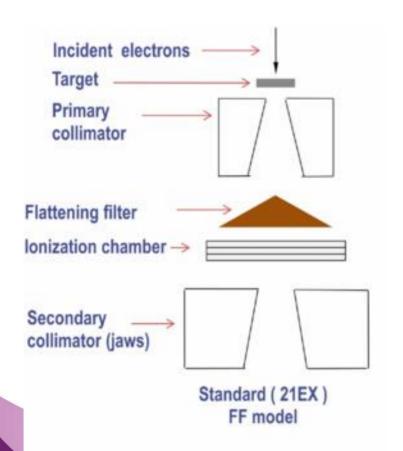
Planning is a collaborative and dynamic process

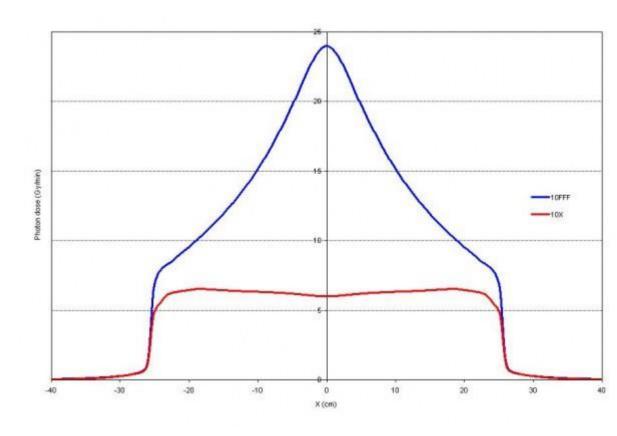




What Has Changed in the World of Planning?

- Flattening Filter Free (FFF)
- Characterized by high dose rate, cone-like fluence profile, increased superficial dose, reduced out-of-field dose





Flattening Filter Free (FFF)

- Not a new concept (1990s) but gained momentum with SRS and SBRT increasing in clinical use
- Rationale and *Theoretical* Benefits
 - Increase dose rate (300-600) to 1400-2400
 - Decreased beam on time (consider even more when combined with VMAT)
 - > Improve efficiency
 - Patient comfort and stability
 - Intrafraction motion
 - Particularly attractive for treatments with high dose per fraction or respiratory motion management (eg DIBH)
 - Lower leaf leakage and out of field dose
 - Possibly offset by an increase in MUs compared to FF beams



What Has Changed in the World of Planning?

- SBRT Planning
- The goal SBRT is to deliver *very* high doses/fx to the target to induce maximum damage
 - > "ablative" doses
- Aim to minimise the volume of healthy tissue receiving a high dose per fraction
 - Dose to OARs is very important due to high dose/fx and increased risk of toxicity
- Traditional dose homogeneity is less of a concern
 - > Up to 160% dose maximum is not uncommon

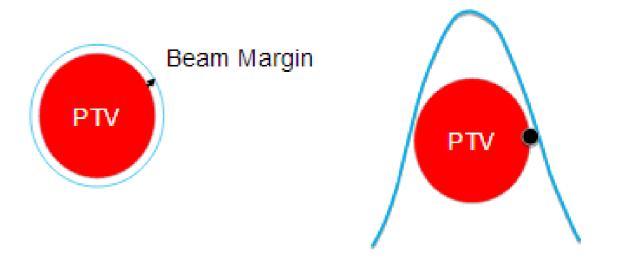


SBRT Planning

Standard approach



SBRT approach





What Has Changed in the World of Planning?

- "Isotoxic" Treatment Planning
- Pioneered by MAASTRO
- Most data for this approach comes from lung cancer with Spinal Cord and MLD as the toxic endpoints
- Moves away from the "one size fits all" approach for dose prescription
 - Dose escalation is based on patient specific OAR DVH results
- Dose escalate the PTV until the OARs reach their tolerance



What Has Changed in the World of Planning?

- Adaptive Radiotherapy
- <u>Definition:</u> "Adaptive radiotherapy involves changes to the radiotherapy plan during treatment on the basis of patient specific observations that were not taken into account during initial planning" (*Gregoire et al., 2012*)
- Incorporates systematic measurements of treatment variations into a closed-loop RT treatment process
- Provides feedback to re-optimise the treatment plan early on during the fractionated course of RT
- Delivers treatment that is customised to the **daily** patient target volumes

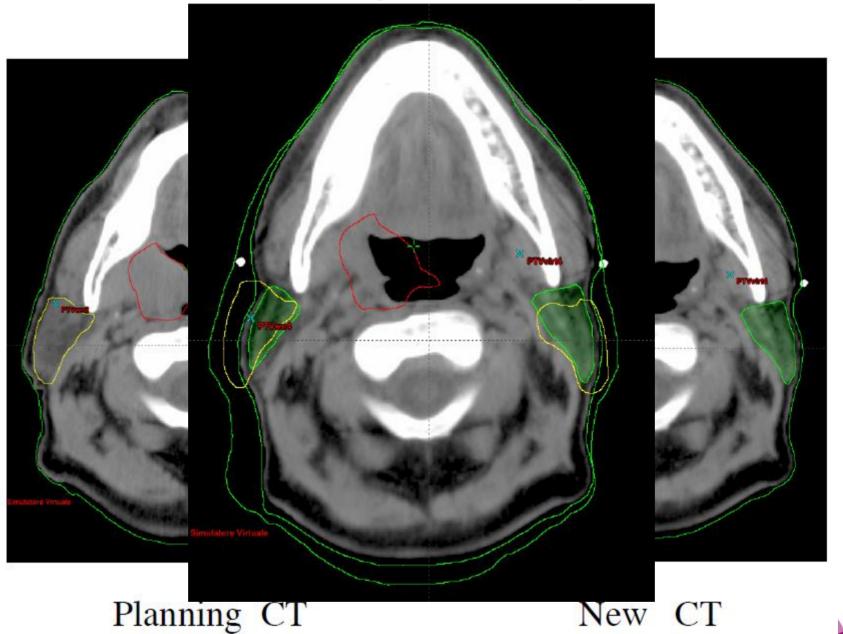
Principles of ART

• Can be adapting to changing geometry or changing geometry and delivered dose

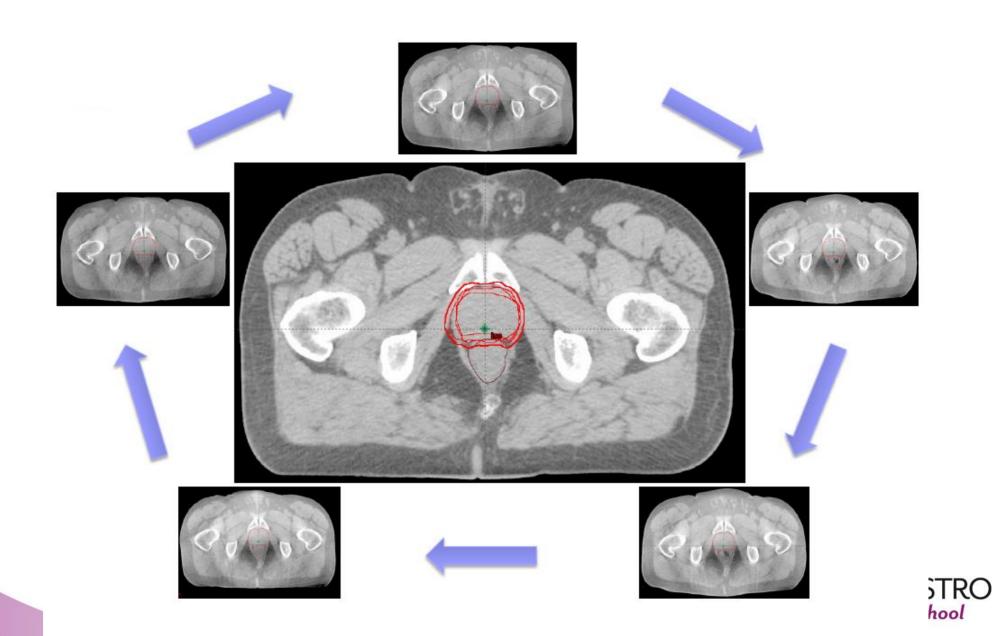
- Approaches:
 - Completely Online
 - Library of Plans
 - > Offline
 - Composite CTV at treatment initiation
 - Scheduled replan
 - Unsceduled replan



Adapting Planning on CT



Adaptive Planning Based on CBCT 1



Adaptive Planning Based on CBCT 2

- Recalculation of planned dose using CBCT
- Are HU on CBCTs accurate?
 - Some conflicting evidence in the literature
 - Depends on the quality of your CBCT
- Options to overcome these uncertainties:
 - Pixel correction technique
 - CT numbers from conventional CT are applied to CBCT
 - Deformable registration
 - Deform planning CT to the CBCT to calculate "dose of the day"
 - This is a move towards "online" ART



Be Careful of Potential Limitations!







Geometric accuracy of bladder plan libraries

- A plan library strategy does not necessarily guarantee geometric accuracy
 - □ Risk of geographical miss due to intrafractional bladder filling in 6 directions (Murthy, 2011)
 - □ No suitable plan in plan libraries (Foroudi, 2011, Lalondrelle 2011, Tuomikoski 2011, Gronborg 2015, Vestergaard 2014)
 - □ Confusion in plan selection (Tuomikoski 2011, Meijer 2012)
 - □ Inappropriate plan selection (Foroudi 2014, Meijer 2012) ESTRO

Logistics of implementation of plan libraries

- ART is not currently feasible for all clinical departments
 - □ Interobserver variability in plan choice posteducation (Kuyumcian et al 2012, Hutton et al 2013)
 - □ Availability of technology may hinder ART implementation (Hutton et al 2013, Murthy et al 2011, Meijer et al 2012)
 - □ Additional education: Cost and Time (MacDonald et al 2013, Lalondrelle 2013, Meijer et al 2012, Wright et al 2008)
 - □ Resource implications on daily workflow (Burridge et al 2006, Hutton et al 2013, Wright et al 2008)



What Has Changed in the World of Planning?

- Automated Planning
- From basic class solution to "Knowledge Based Planning"







What Will Planning Look Like in the Future?

- Will continue to increase in complexity
 - Biological optimisation
 - ART and personalised approach based on Radiomics based analysis of pre treatment and during treatment imaging
- *Radiomics* is the extraction of quantitative imaging features that can be combined with clinical data
- Will move from a separate planning room to the linac
 - MRI linac
 - Online reoptimisation
 - Online ART



Take Home Message

- Have an awareness of what to expect from your plan
- If you don't get that, always ask why?
 - Having an understanding of why the dose has behaved that way will help you find a solution to the problem
- Be guided by the literature
 - ➤ Almost all dosimetry papers will outline their planning process
 - Beam arrangement
 - Energy
 - Prescription method used
 - Critical analysis is needed!



Take Home Message

When reading the literature, read carefully!

Good, that sounds like ICRU 50

Methodology:

"The radiation dose was prescribed to the PTV, such that 100% of the PTV received >95% of the prescribed dose and that no region in the field received greater than 107% of the prescribed dose"

• Results:

Hang on a minute?!

"All treatment plans showed adequate coverage of the way wolume, with more than 95% of volume of PTV1 and PTV2 receiving greater than 95% of the prescribed dose."



In-room imaging modalities

Martijn Kamphuis MSc MBA Research Radiation Therapist IGRT

> Department of Radiotherapy Amsterdam, the Netherlands



Content of the presentation

•Why do we need imaging on the linac?

Imaging modalities

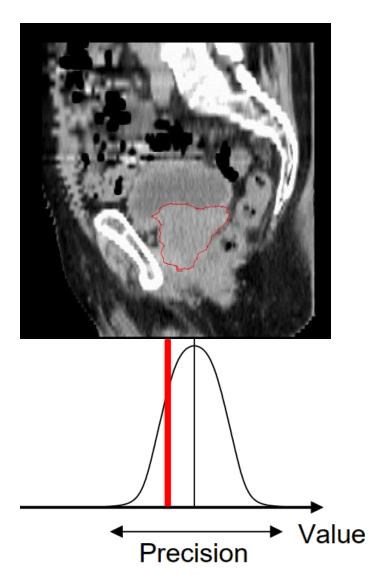
- ➤ How do the work?
- What can we do with them?
- > Pros and cons





At the start of treatment

- Single CT introduces systematic errors*:
 - Delineation errors
 - Organ position and shape at time of localization
 - Phantom transfer errors
 - o Geometric imaging error
 - o Treatment planning system error
 - o Linear accelerator geometry error
 - Set-up error at time of localization
 - TPS beam algorithm error
 - Breathing positional error



*McKenzie et al., BIR 2003 Image courtesy: Marcel van Herk

In fact...it's just a snapshot



Why do we need imaging on the linac?

To reduce systematic and random geometrical errors

- Monitor/adapt to patient anatomy/pathology
 - Plan of the day
 - > (Ad hoc) replanning
- To document the treatment accuracy
 - Margin calculation
 - Incident analyses



Imaging modalities

- Ultrasound systems
- Electromagnetic tracking
- Portal Imaging (EPID)
- kV cone beam CT
- 3D CBCT
- MV (CB)CT

- Surface scanning
- MR linac



Polling: Who is using what?

A.Ultrasound

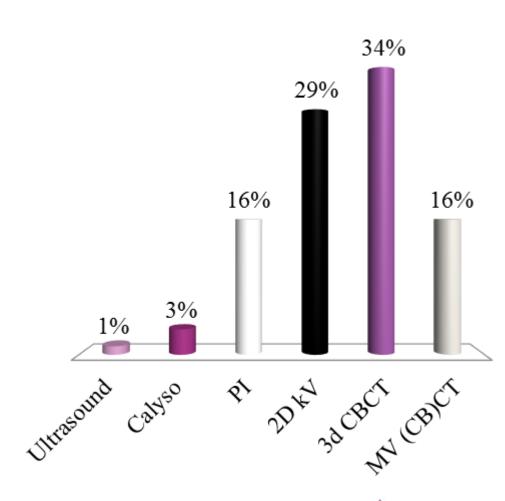
B.Calyso

C.PI

D.2D kV

E.3d CBCT

F.MV (CB)CT





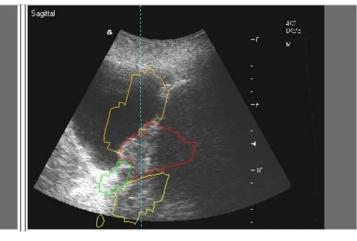


Ultrasound systems

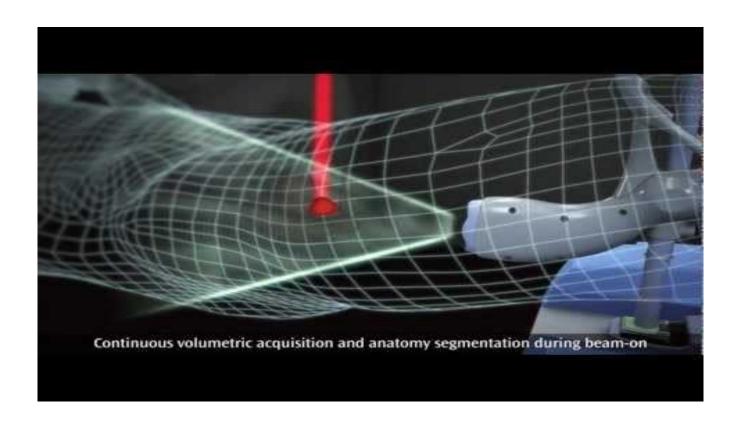
With probe define position target

Infrared enables correlation with linac





More recent developments



https://www.elekta.com/software-solutions/treatment-management/imaging/clarity/?utm_source=clarity&utm_medium=redirect&utm_campaign=redirects

Ultrasound system

Pros:

- Non invasive
- No imaging dose
- (Intra fraction imaging)

Cons:

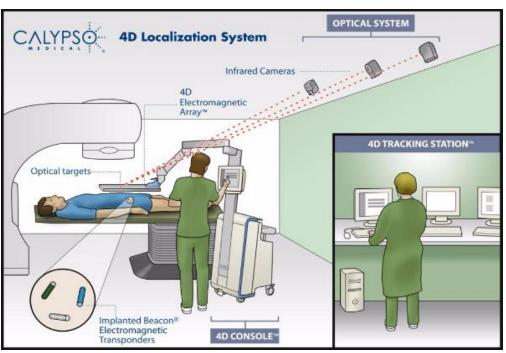
- (User dependent accuracy)
- (No intra fraction information)
- Limited number of indications
 - Prostate
 - Upper abdominals
- (Probes influences position target)
 - Systematic error



Electromagnetic tracking

- Uses implanted fiducials
- Lower magnetic field
- Transponder emits RF









Electromagnetic tracking

Calypso System

The Calypso System allows for real-time tracking of tumors during prostate cancer radiation therapy treatment.

The Calypso System helps doctors track the exact location of a prostate tumor DURING the actual radiation treatment for prostate cancer.



Electromagnetic tracking (GPS)

Pros:

- Continuous real time measurements (10Hz)
 - > Intra fraction monitoring is used for others sites as well
- Non ionizing

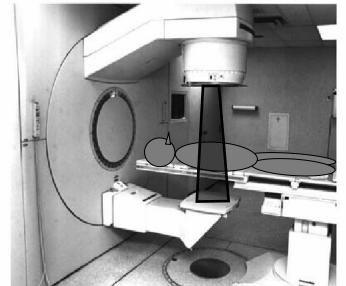
Cons:

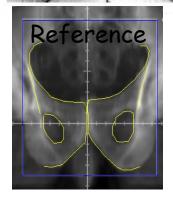
- Limited number of indications
 - Mostly prostate
 - Lung
 - Breast (PBI)
 - Pancreas
- No anatomical information
- Invasive pre imaging procedure



Portal Imaging - physics

- An imager used to detects the photons that cross the patient
- The portal image is compared to a reference image









Goals of Portal Imaging

- Position verification
- Documentation of treatment
- Portal dosimetry (in-vivo)
- QA (MLC adjustment)



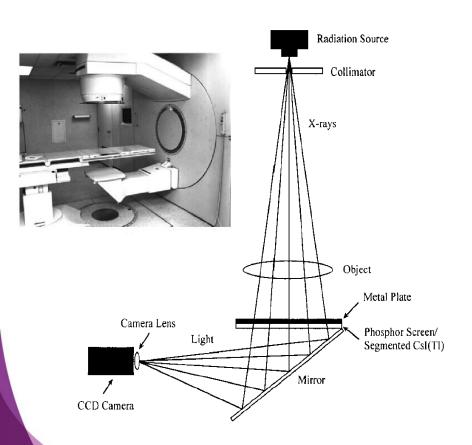


Technical aspects of EPIDs

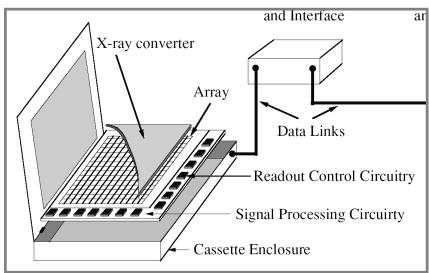
Camera-mirror based systems

Active matrix flat panel imagers (AMFPI)

• also called amorphous silicon imagers







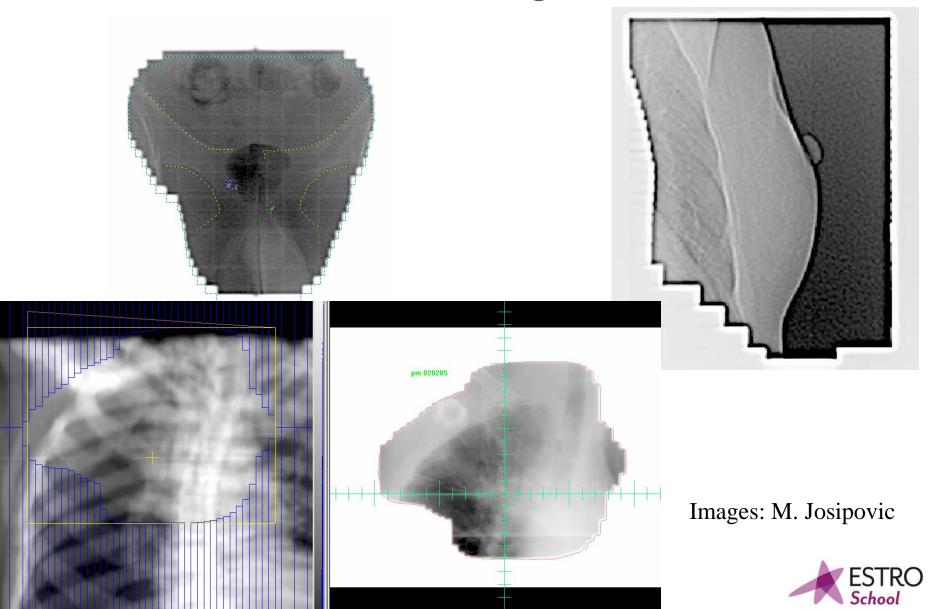
Examples of portal images (open field)



Images: M. Josipovic



EPID – field images



Electronic Portal Imaging

Pros

- Image made with treatment beam
- Imaging during treatment
- Possible to perform dosimetry

Cons

- Surrogate imaging
 - Additional margins
- Imaging dose
 - > Although it is possible to compensate for
- Imaging quality



2D kV imaging

kV source & detector panel

Different approaches:

- kV source moutend on linac
- kV sources on fixed position in room

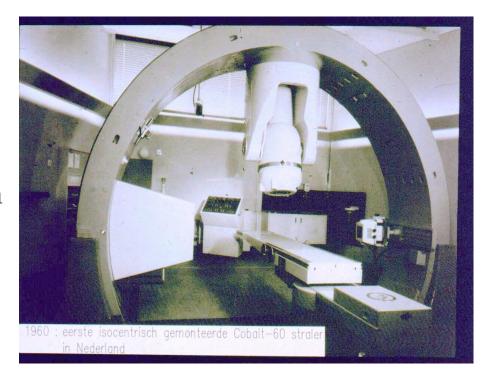
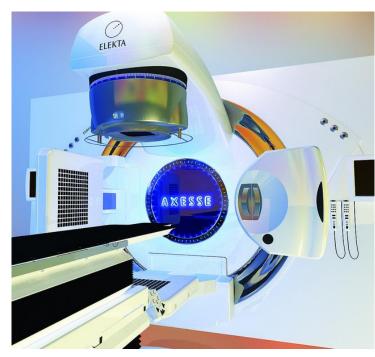


Image: Ben Mijnheer (NKI)



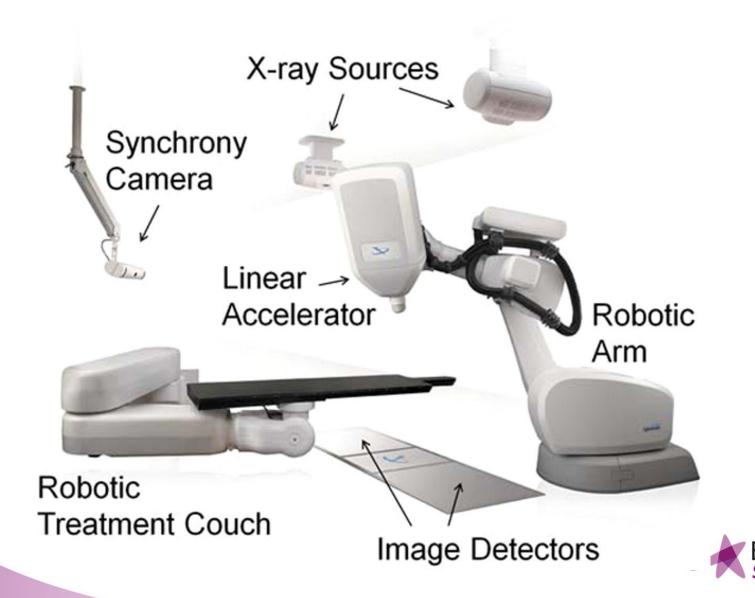
kV source moutend on linac



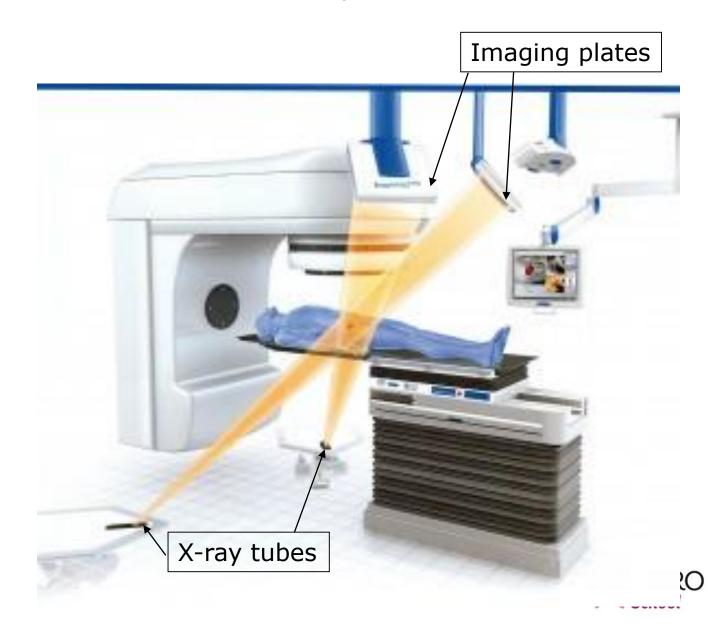




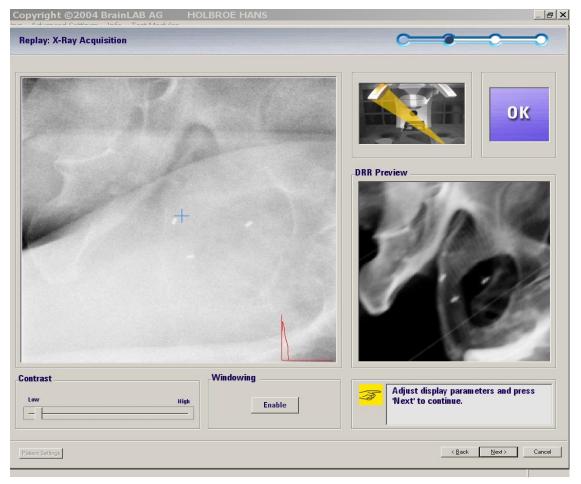
kV imaging: Cyberknife



Exac Trac® IGRT system



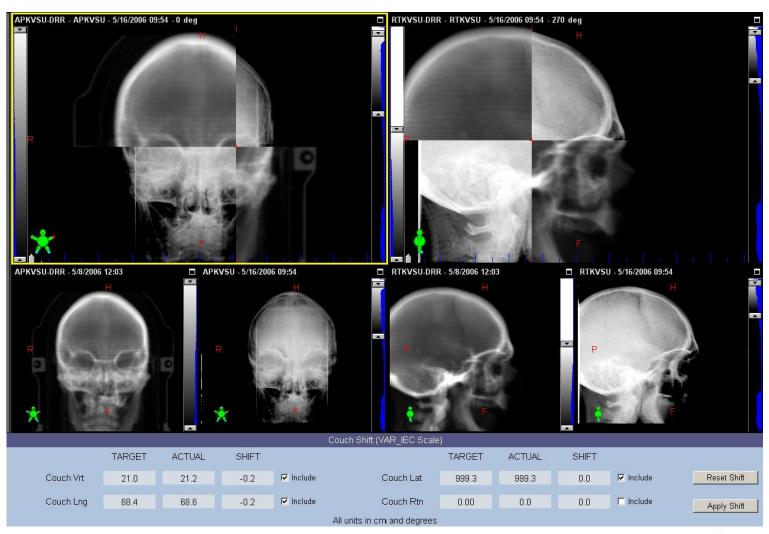
Exac Trac® IGRT system



Images: M.Josipovic



OBI kV imaging



Images: M.Josipovic



kV imaging

Pros:

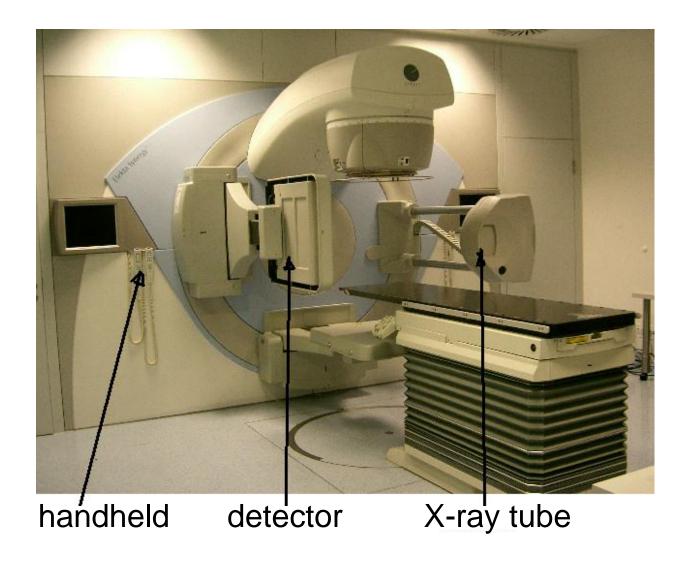
- Imaging dose is low
- High 2D imaging quality
- Real time imaging in some systems (all angles)

Cons

- Limited anatomical information
- In most times it is a surrogate
- Oblique images are difficult to interpret.



Cone beam CT

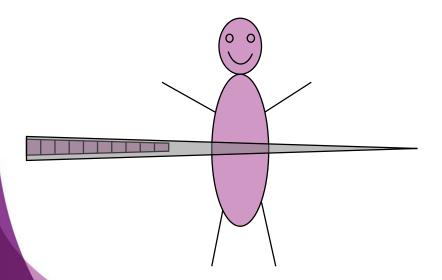




CBCT Acquisition

Conventional CT

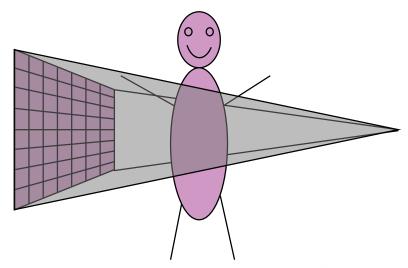
- 'Fan' beam
- 1D detector
- 1 rotation = 1 slice



Courtesy: Peter Remeijer

Cone-beam CT

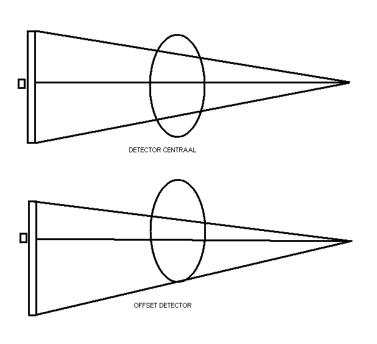
- 'Cone' beam
- 2D detector
- 1 rotation = volume (many slices)





How does it work?

Variable detector position



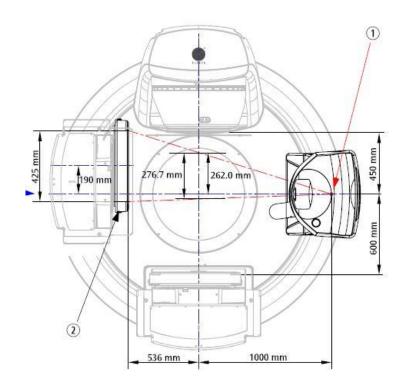




Image registration: Defining the ROI

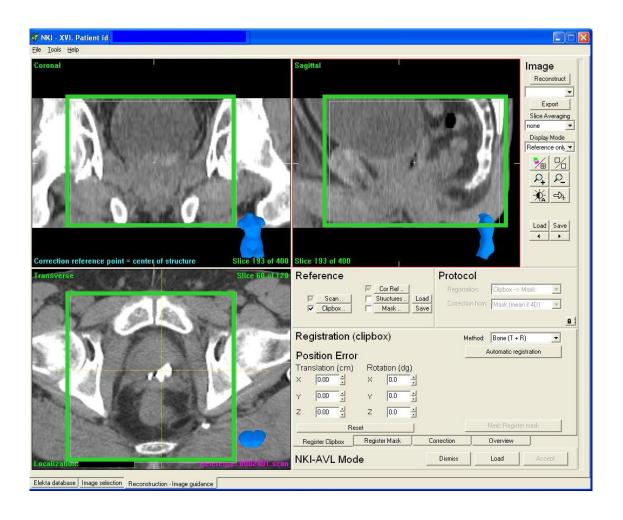




Image registration: Defining the ROI



Image registration

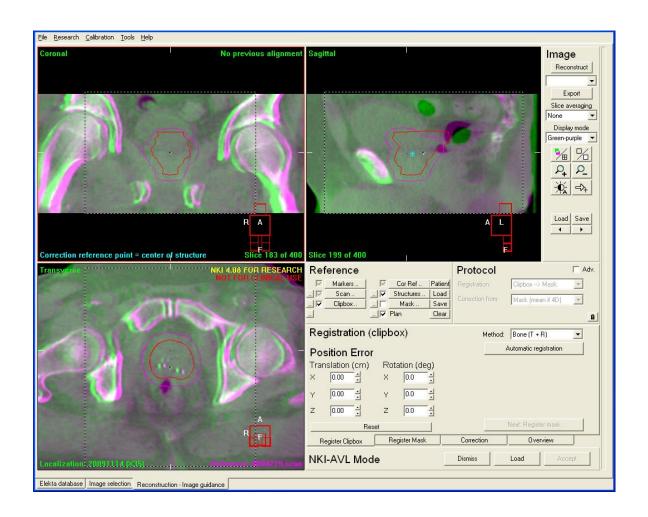




Image registration: bony anatomy

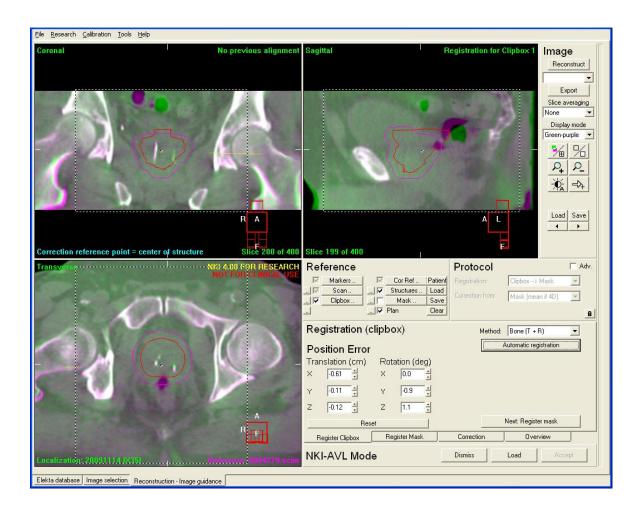
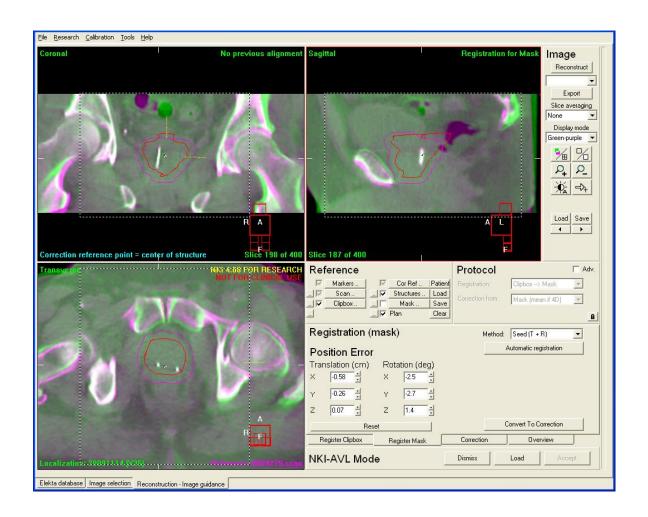




Image registration: fiducial markers





CBCT imaging

Pros:

- Anatomical information
- Imaging dose can be low
- Relatively high imaging quality
- Good to excellent registration algoritms

Cons

- Imaging dose can be substantial
- No real time imaging in some systems
 - Inline scanning still leads to retrospective analyses



MV-(CB)CT

Using:

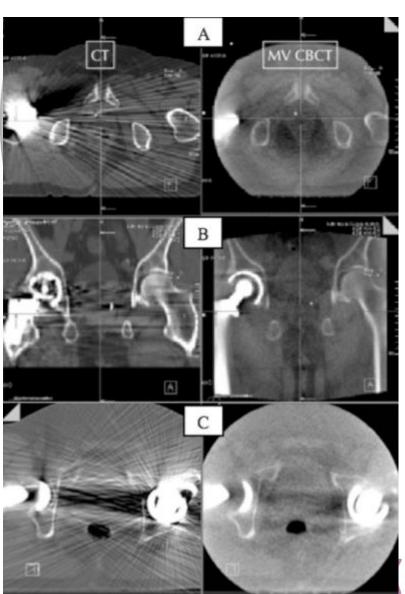
- > Treatment beam
- > Flat panel
- ➤ 3D acquisition

MV-CT:

- > Helical acquisition
- > TomoTherapy

MV-CBCT:

- > 360 degrees acq.
- Siemens Oncor





MV-CT

Pros:

- Anatomical information
- Limited influence of high densities (prostheses)
- Image of the actual absorbed dose

Cons

- Image quality not as good as kV CBCT
- Imaging dose
- Only available as Siemens



Videosystems

Different approaches:

- Infrared tracking of external markers
- Surface scanning

What can you do with these systems?

- Set-up aid
- More important: monitor the patient during treatment:
 - 1. Passive: monitoring set-up accuracy
 - 2. Active: correlate motion with treatment (e.g. gating or DIBH)



Exac trac infrared

- Infrared marker,
 - placed on fixed spots
- Tracking of the markers during RT
 - Correlate with respiration (tracking/gating)



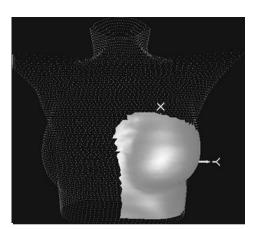


Images: M.Josipovic

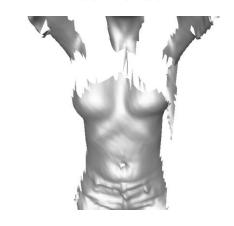


Surface scanning





Left side



Images: T.Alderliesten



Infra red systems

Pros

- No imaging dose
- Enables tracking and gating
- Real time measurements
- Surface scanning:
 - Pre treatment set-up check

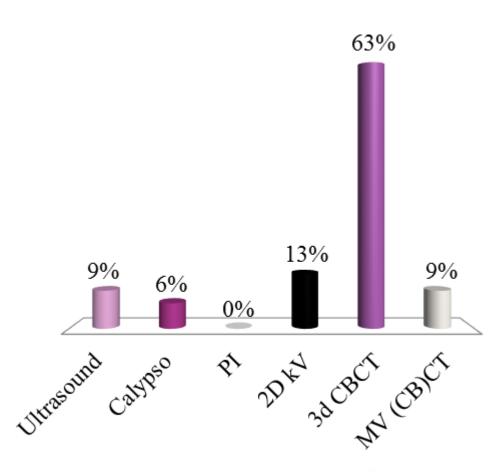
Cons

- It's an aid
 - Can never be a stand alone system
- Surrogate



Which one do you prefer most in prostate?

- A. Ultrasound
- B. Calypso
- C. PI
- D. 2D kV
- E. 3d CBCT
- F. MV (CB)CT





Which one do you prefer most in lung?

A.Ultrasound

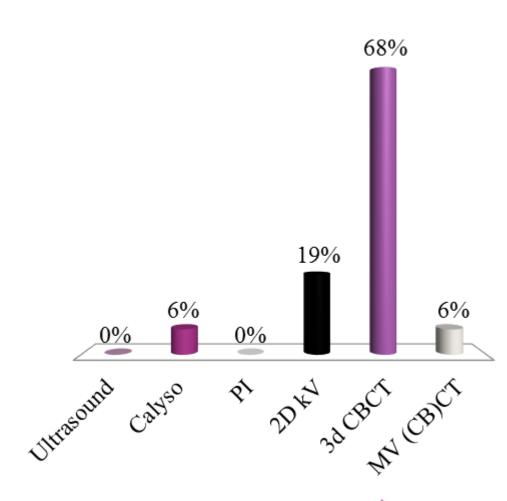
B.Calyso

C.PI

D.2D kV

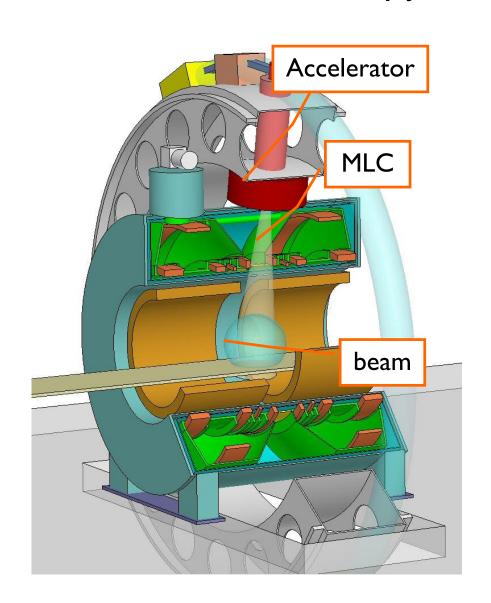
E.3d CBCT

F.MV (CB)CT





Integrating MRI functionality with external beam radiotherapy



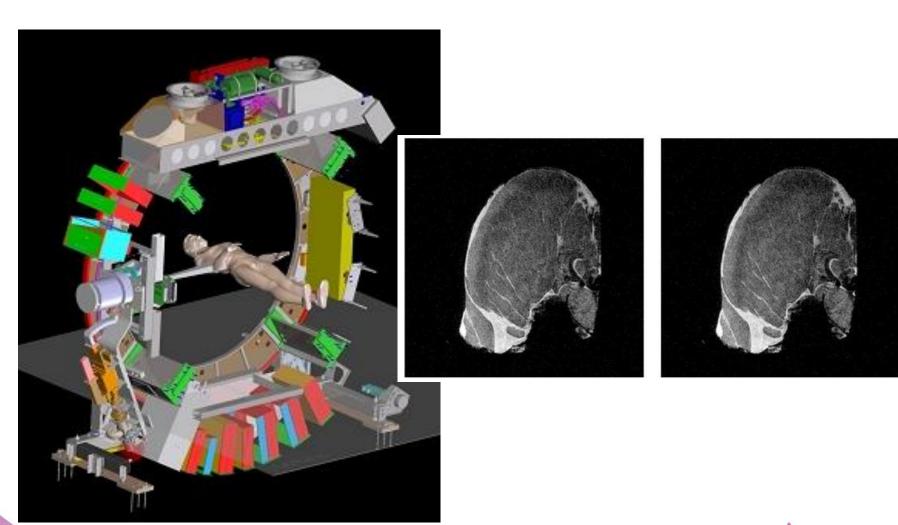


Integrating MRI functionality with external beam radiotherapy





Gantry design MRL: (MRI-Linac)





MRIdian: MR Cobalt





MR linac

Pros

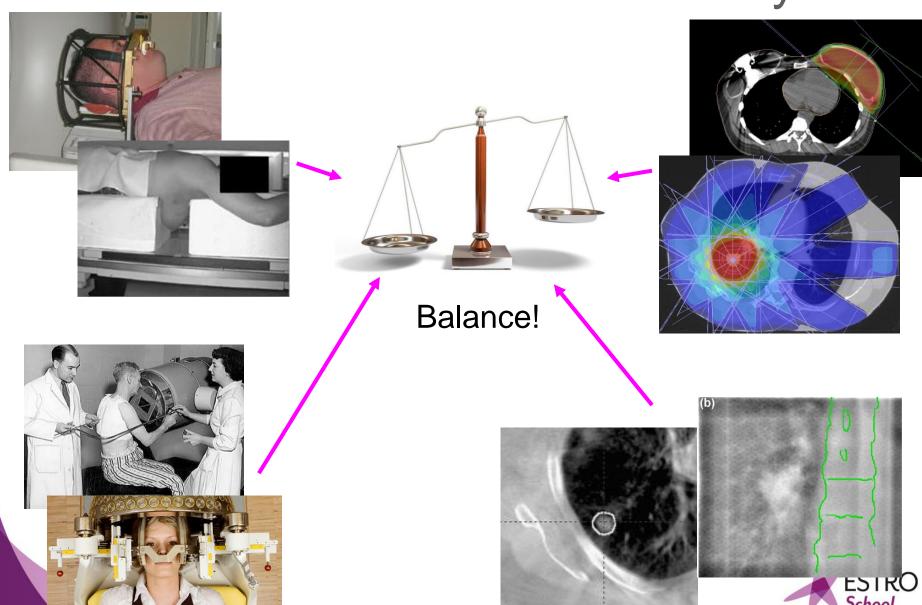
- Optimal image quality
- Intra fraction imaging

Cons

- MR-Linac:
 - quite expensive
 - Under development, mainly research
- Cobalt treatment: linac upcoming
- Challenging Treatment planning (1,5 Tesla)
 - Secondary electrons are influenced by the magnetic field



How accurate should the delivery be?



Clinical rationale for image-guided radiation therapy (IGRT)





Jose Lopez, M.D., Ph.D Radiation Oncology University Hospital Virgen del Rocio Seville, Spain

Advanced skills in modern radiotherapy

Prague, Czech Republic –11-15 June 2016

Learning Objectives (IGRT)

- Learn the clinical rationale for IGRT
 Why we should do it
- Learn the challenges in achieving precision and accuracy
- Understand the benefits and limitations of IGRT
- Learn the evidence that supports the use of IGRT



Peterpedia



Wikipedia

- IGRT is the process of frequent <u>two and three-dimensional</u> <u>imaging</u>, during a course of RT, <u>utilizing the imaging</u> coordinates.
- The patient is localized in the treatment room in the same position as planned from the reference imaging dataset.
- An example of IGRT would include:
- localization of a **CBCT** dataset with the planning **CT** dataset
- matching planar kV or MV images with DRRs



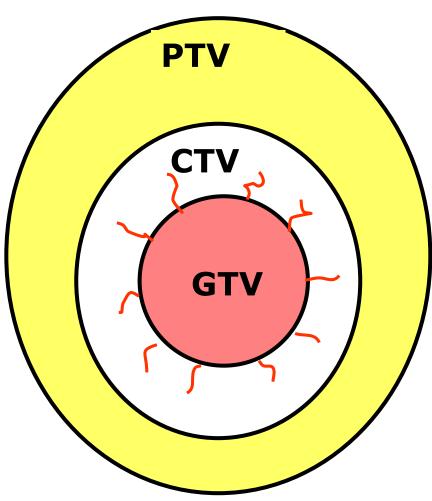
Why do we need IGRT?





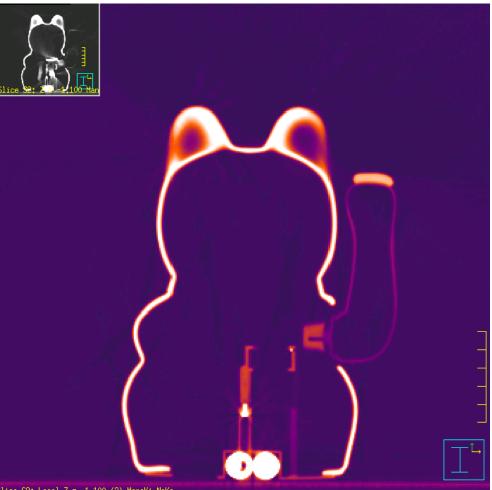
Tumor motion



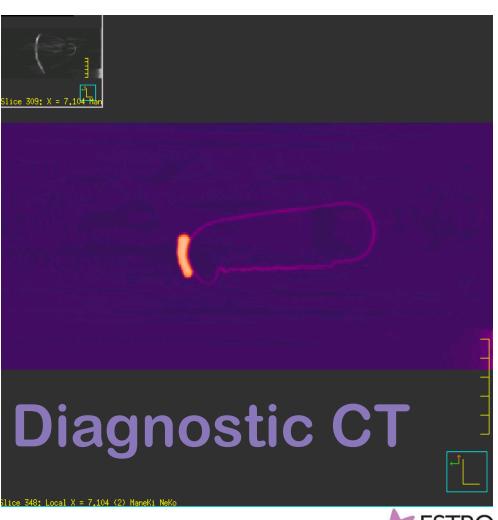








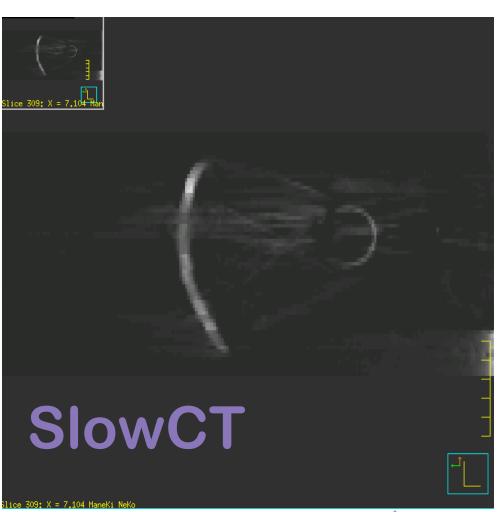




Courtesy Santiago Velazquez





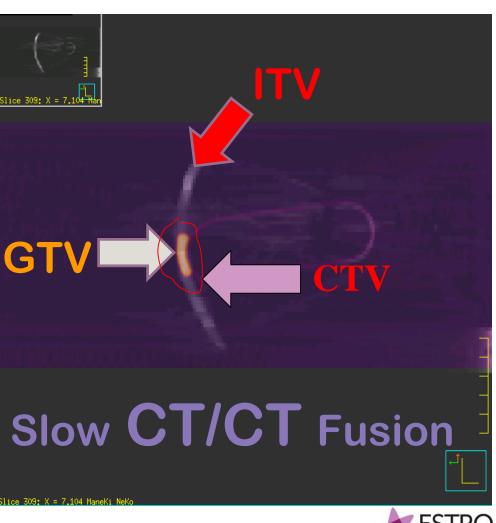


Courtesy Santiago Velazquez



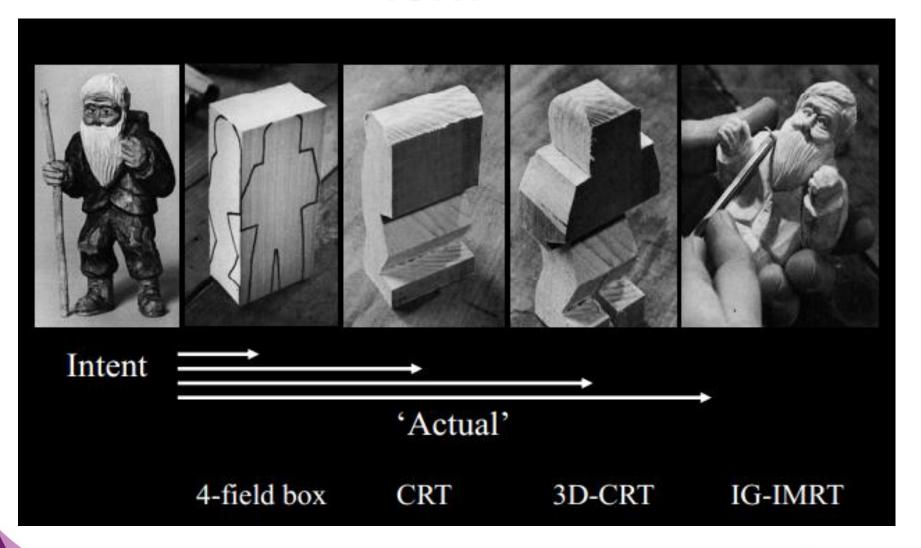
SlowCT/CT Fusion



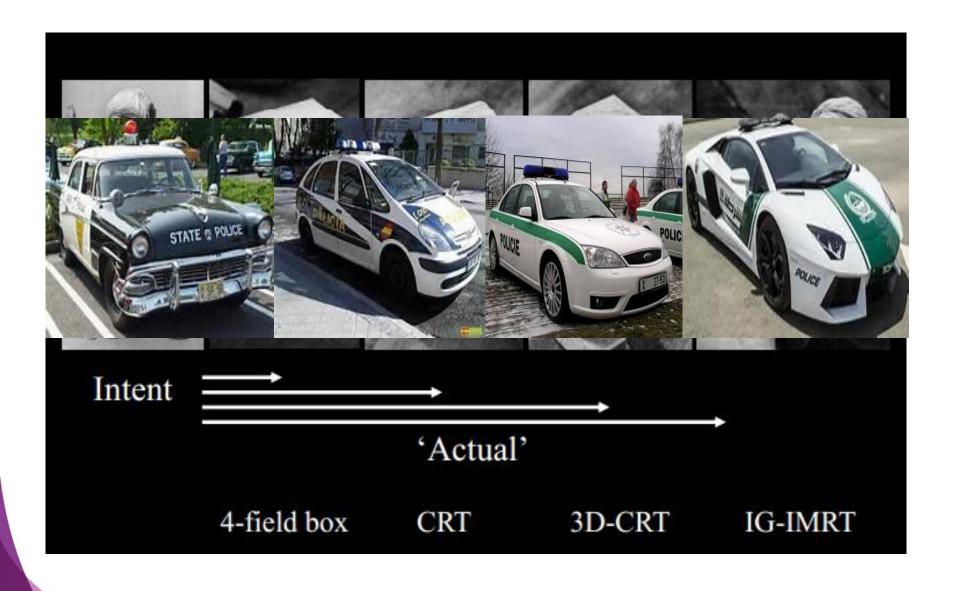


Courtesy Santiago Velazquez

IGRT





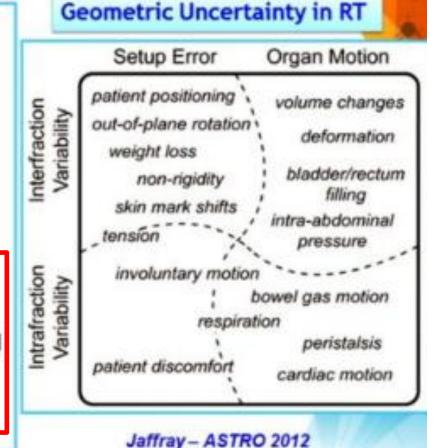




Rationale for IGRT

Quality of Radiotherapy Delivery

- Quality of radiotherapy (RT) delivery is one of the important determinants of patient outcomes
- Efforts to improve the quality of RT delivery include:
 - Accurate target delineation
 - Robust plan optimization
 - Minimizing day-to-day setup variation
 - Tracking intra-fractional organ motion
- Monitoring & adapting to inter-fractional tumor and normal tissue changes
- Enhance deliverability (spatial access)

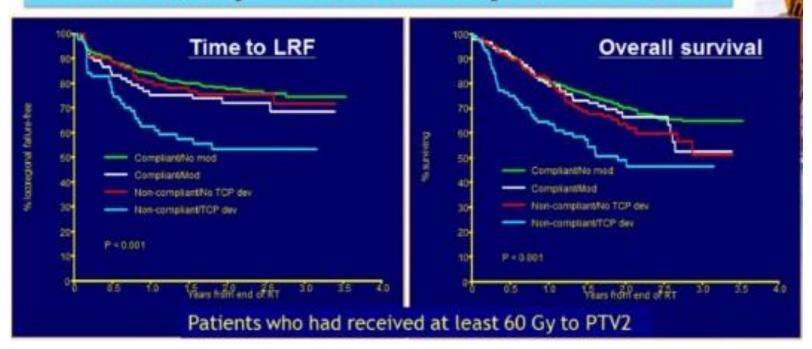


- >800 patients across four Continents
- Influence by accrual numbers

Critical Impact of Radiotherapy Protocol Compliance and Quality in the Treatment of Advanced Head and Neck Cancer: Results From TROG 02.02

Leuer J. Piners, Brian O'Sullivan, Jordi Giralt, Thomas J. Fietgerald, Andy Trom, Jacques Bernier, Jean Bosobio, Kally Yson, Richard Fisher, and Danny Rischin

Radiation Quality Matters: Results by deviation status



Imaging for treatment verification

1980's – port films

1990's - emergence of MV portal imagers

in-room ultrasound localization

marker-based localization

Fluoroscopic tracking

2000's – flat panel imaging

KV digital imaging

CBCT

MV CBCT

CT "on rails"

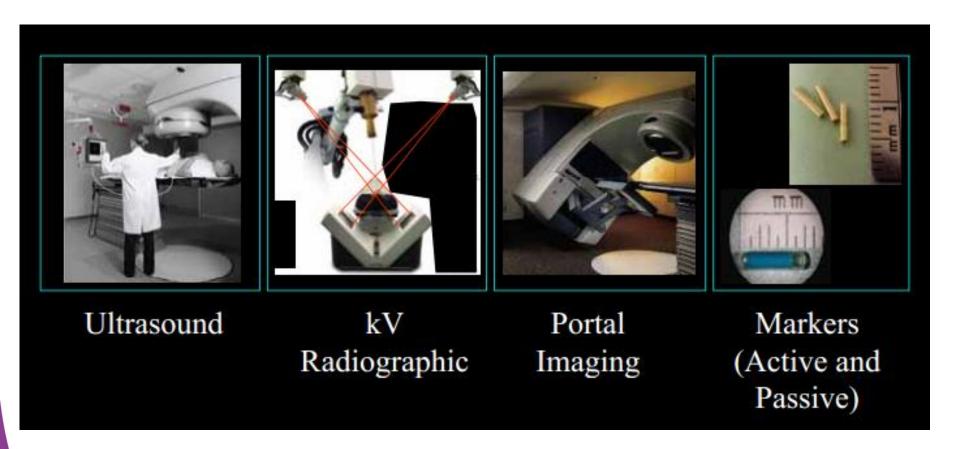
Emerging - Electromagnetic localization and tracking

surface tracking

in-room MRI

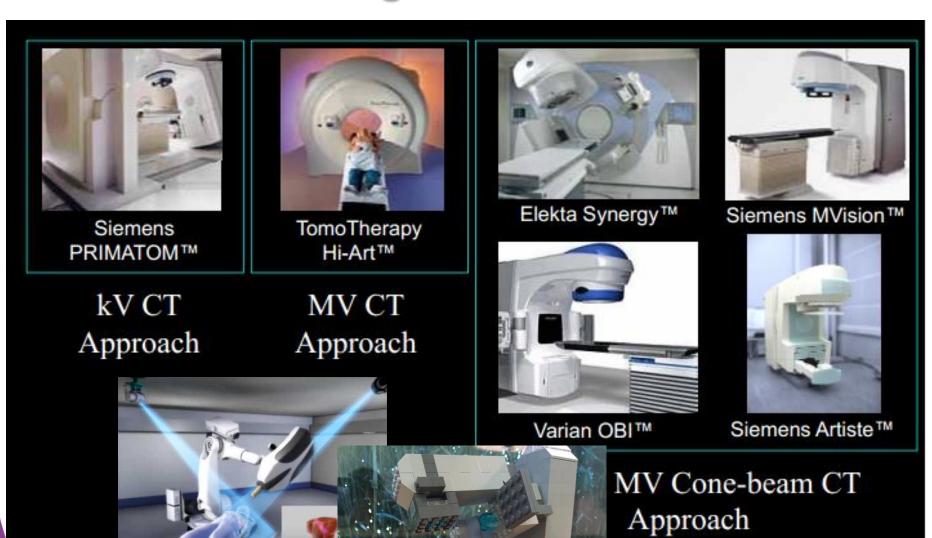


Technologies available for IGRT





Technologies available for IGRT



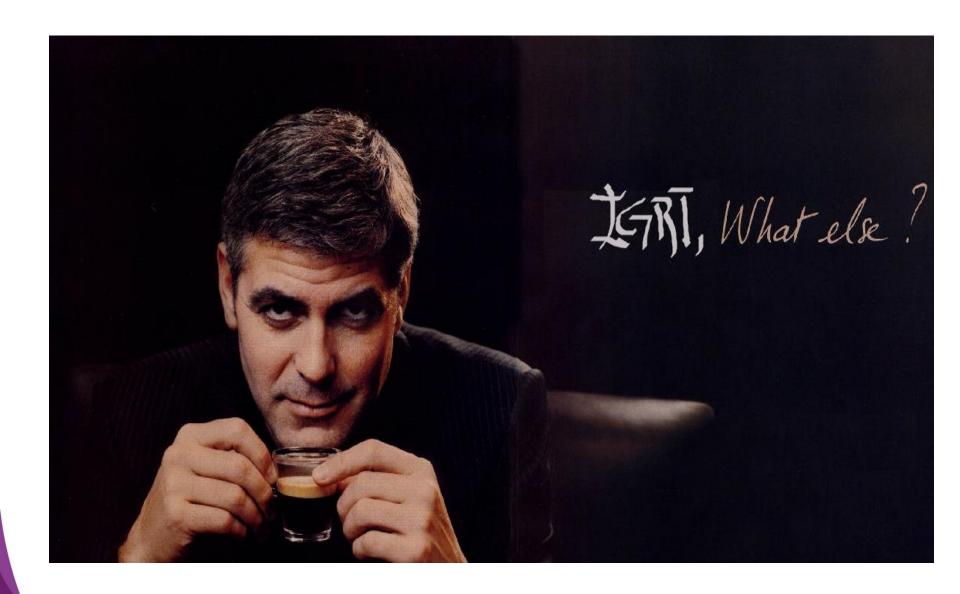


Adoption of new RT Technology

- Vendor and developer motivation
- Healthcare provider's incentive
- Patient and their family's perception
- Public health provider and Policy maker's concern
- Adoption of these techniques is often hasty

Mainly focus on technological capacity rather than evidence-based sptepwise approach







Rising Cost of Radiotherapy



Contents into postate at followsr framedised

Radiotherapy and Oncology

(market barringlega; www.htmpscoopsarket.com



Does of codeotherapy:

The cost of vadiotherapy in a decade of technology evolution

Everyo Van de Werf** Jan Verstrame*, Volande Lievenn**

Table 2
Comparison of the costs at the activity-group level for respectively 2000 and 2009.

	2000 (€)	%	2009 (€)	%
Treatment preparation	839,247	31	1,480,112	27
First patient contact	137,066 423,162 25,540 253,479	5 16 1 9	197,612 667,324 114,611 500,565	4 12 2 9
Simulation				
Delineation				
Dose calculation				
Treatment delivery	1,872,695	69	4,059,947	73
Quality assurance (QA)	283,192	10	1,252,789	23
General at start	40,561	1	102,459	2
Patient specific	69,606 60,324	0 3 2	48,678 68,519 791,248	\searrow_4^1
Supervision plan				
Portal imaging				
In vivo dosimetry	32,423	1	76,064	1
Chart round	80,278	3	165,821	3
Daily radiotherapy delivery	1,508,306 44,820	56 2	2,501,649 116,816	45 2
Clinical follow up				
Discharge	36,377	1	188,693	3
Total	2,711,942	100	5,540,059	100



Rationale for IGRT

Adoption of new IGRT techniques should be based on clinical rationale/evidence and clinical needs:

- What is the clinical evidence for the claims
- ➤ It is better/lower cost than current standard
- > It can tackle a currently unsolvable clinical problem
- > It is any purported benefit
- What is the clinical indication
- What are the limitations/risks
- Do we have resources and demand
- CLAIM IS NOT EVIDENCE!!



Challenges in RT delivery:

- Day-to-day setup variation
- Intra-fractional organ motion
- Inter-fractional tumor and normal tissue deformation

IGRT Claims: (compared to conventional 3D Conformal)

- IGRT improves the precision and accuracy of RT delivery
 - Minimizes day-to-day setup variation with daily imageguidance using appropriate matching surrogate
 - Tracks intra-fractional organ motion
 - Tracks inter-fractional tissue deformation (adaptive)
- Does it translate into better clinical outcomes?





Radiotherapy and Oncology 78 (2006) 119-122 www.thegreenjournal.com

Special commentary

From IMRT to IGRT: Frontierland or Neverland?

C. Clifton Ling^{a,*}, Ellen Yorke^a, Zvi Fuks^b

^aDepartment of Medical Physics, and ^bDepartment of Radiation Oncology, Memorial Sloan Kettering Cancer Center, New York, NY, USA

Abstract

The recent enthusiasm for real-time image guidance in radiotherapy (IGRT) is in part due to the commercial availability of advanced on-line imaging technologies. Perhaps more important than its potential to improve conventional radiotherapy, IGRT may lead to a paradigm shift in facilitating hypo-fractionated or single-dose treatment. However, there are uncertainty regarding features and approaches of competing IGRT systems and as to whether a sub-set of the features of an ideal IGRT system would suffice for specific disease sites and clinical applications. Clinical studies are necessary for the quantification of benefit needed for evidence-based medicine (Bentzen, SM. Radiation therapy: intensity modulated, image guided, biologically optimized and evidence based. Radiat Oncol 2005;77:227-230).

© 2005 Elsevier Ireland Ltd. All rights reserved. Radiotherapy and Oncology 78 (2006) 19-122.



IGRT





EDITORIAL

Will IGRT live up to its promise?

MARCEL VAN HERK

The Netherlands Cancer Institute/Antoni van Leeuwenhoek Hospital, Amsterdam, the Netherlands



Evidence levels

Levels		Type of Evidence		
I 1A 1B	1A	Systemic review (with homogeneity) of RCTs		
	1B	Individual RCT (with narrow confidence intervals)		
	10	All or none study		
	2A	Systematic review (with homogeneity) of cohort studies		
	2B	Individual cohort study (including low quality RCT, e.g. <80% follow-up)		
2C		"Outcomes" research: Ecological studies		
	ЗА	Systematic review (with homogeneity) of case-control study		
	3B	Individual Case-control study		
IV	4	Case series (and poor quality cohort and case- control study)		
٧	5	Expert opinion without explicit critical appraisal or based on physiology bench research or "first principles"		



Guyatt et al. JAMA 2000



Improved Clinical Outcomes With High-Dose Image Guided Radiotherapy Compared With Non-IGRT for the Treatment of Clinically Localized Prostate Cancer

Michael J. Zelefsky, M.D., "Marisa Kollmeier, M.D., Brett Cox, M.D., "Anthony Fidaleo, B.A., "Dahlia Sperling, B.A., "Xin Pei, Ph.D., "Brett Carver, M.D., Ph.D., Jonathan Coleman, M.D., Michael Lovelock, Ph.D., and Margie Hunt, B.S.

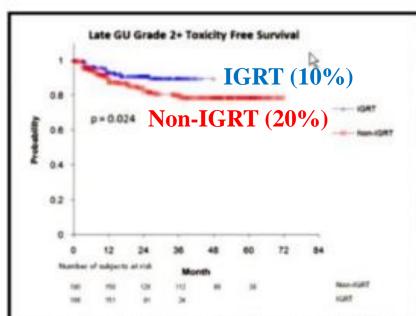


Fig. 1. Comparison of actuarial likelihood of grade 2 or higher late urinary toxicity for patients treated with image-guided radiotherapy (IGRT) to 86.4 Gy vs. intensity-modulated radiotherapy.

IGRT (daily imaging with fiducial marker) vs. non-IGRT (weekly imaging without marker):

- √ Same Rx dose: 86.4 Gy
- √ Same PTV margin
- ✓ Similar delivery technique: IMRT

IGRT vs non-IGRT

- Lower G2+ GU toxicity (3-y: 10 vs 20%)
- Higher biochemical tumor control in highrisk cohort
- Limitation:
- √ Retrospective review
- √Slightly different study period
 - Non-IGRT: 2006-2008
 - IGRT: 2007-2009
- ✓ Short follow-up: median 2.8 years, esp. IGRT cohort

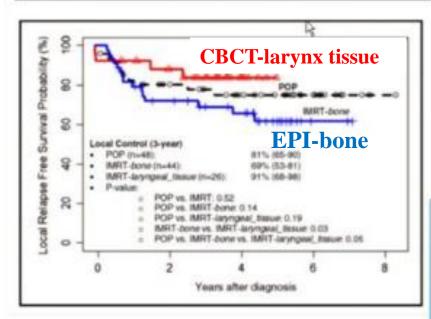
IGRT reduced daily setup variation ⇒ better clinical outcomes (level 3B evidence)



CBCT (vs. MV-EPID): better soft tissue visualization

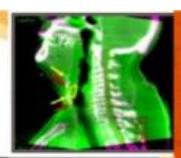
Outcome following IMRT for T2 glottic cancer: the potential impact of image-guidance protocols on local control

Albert Tiong - Shao Hai Huang - Brian O'Sullivan - Indranil Mallick - John Kim-Laura A. Dowson - John Cho - Jolie Bingash - Andrew Bayley - Andrew Hope -Engene Yn - Stephen Breun - Andrea McNiven - Raiph Gilbert - Wei Xu - John Waidron



Level 4 evidence





IMRT (CBCT to laryngeal tissue) vs IMRT (MV-EPID to bone)

- Higher LC (97 vs 69% at 3-yr)
- Limitations:
- ✓ Retrospective review
- ✓ Different study period
 - Non-IGRT: 2006-2008
 - IGRT: 2008-2010
- · Smaller sample size (44 vs 26 patients)
- IMRT with MV-EPID matching to bone could result in geographic miss
- IGRT with CBCT allows selecting appropriate matching surrogate (laryngeal soft tissue) ⇒ improved setup accuracy ⇒ improved clinical outcomes

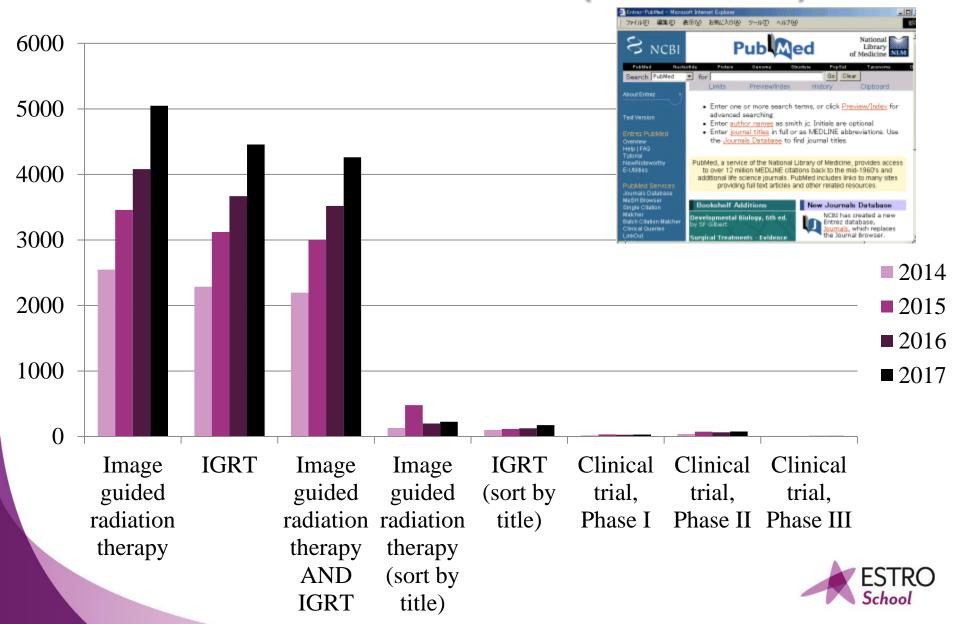


Sources for Clinical Rationale

- Premarketing research work:
 - Randomized trial is ideal but often lacking
 - Difficult to conduct (small "window of opportunity")
- Published literature
 - Various quality: high level of evidence is scant
 - Be aware of publication bias, reporting bias, reviewers' bias, omission bias
- Official and unofficial communication
 - Conference, courses, symposium, expert narration
 - Subject to bias, especially vendor sponsored symposium
- Own institutional experience following implementation
 - Cumulative, prospective, and reflective
 - Requires close monitoring and timely feedback



Cites in Pubmed (2014-2017)



Clinical trial, Phase III (compare new treatments with the standard)

- Standard-dose (60Gy) vs high-dose RT (74Gy) for NSCLC patients (RTOG 0617): negative results
- > RT planning was more likely to be noncompliant in the high-dose group (26% vs 17%, P = .02)
- ➤ They used both 3D and IMRT
- No details about IGRT
- RTOG 0631 phase 2/3 study of image guided stereotactic radiosurgery for localized (1-3) <u>spine</u> metastases: <u>phase 2 results</u>.
- A randomized hypofractionation <u>dose escalation</u> trial for high risk <u>prostate</u> cancer patients: interim analysis of acute toxicity and quality of life in 124 patients.

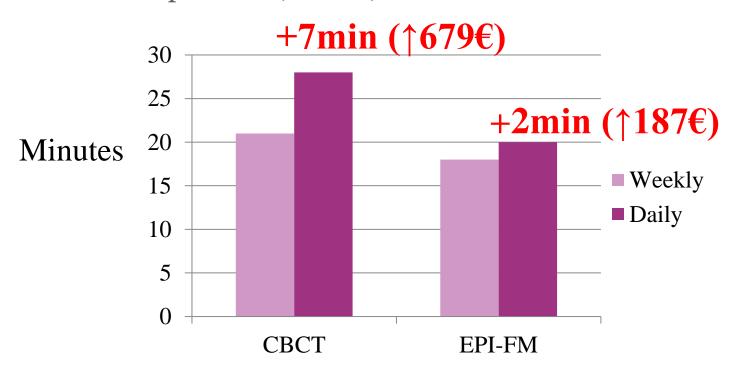
Clinical trial, Phase III

- **Cost** of prostate IGRT: results of a randomized trial.
- Prognostic **impact of abdominal adiposity** in prostate patients.
- **Recommendations** for implementing **SBRT** in peripheral stage IA NSCLC: phase III ROSEL study.
- **Dosimetric experience** with accelerated partial breast irradiation using image-guided interstitial brachytherapy.



COST and TIME

• N=208 patients (France)

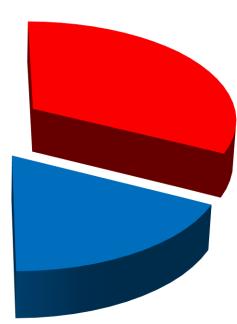


• The incremental costs due to different IGRT strategies are relatively **moderate**.



Clinical trial, Phase II (N = 74) (if a new treatment works well)





- Prostate (24)
- Lung (13)
- Oligometastases (5)
- **■** Liver (4)
- Head and Neck (5)
- Rectum (3)
- Soft tissue sarcoma (1)
- **■** Breast (5)
- Cervix (2)
- Pancreas (3)
- Spinal metastases (3)
- Esophagus (1)
- Gastric (1)



Toxicity

Grade 0	None
Grade 1	Mild
Grade 2	Moderete
Grade 3	Severe
Grade 4	Intensive care
Grade 5	Fatal

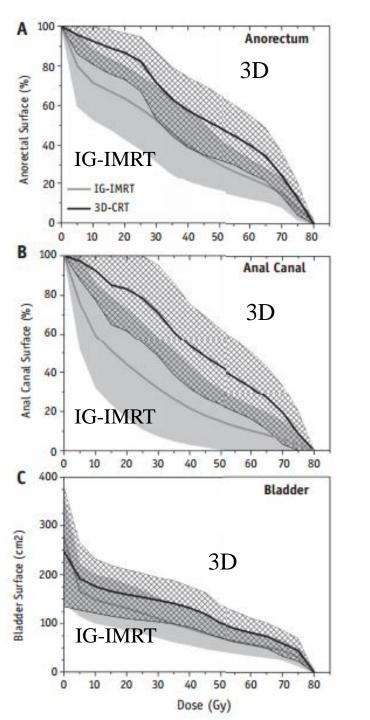


Phase II (Prostate)

Variable	3D-CRT $(n=215)$	IG-IMRT (n=26
Mean (±SD), y	68.9 (±6.3%)	70.5 (±6.0%)
T category		
1	36 (16.7%)	40 (15.4%)
2	97 (45.1%)	89 (34.2%)
3a	53 (24.7%)	102 (39.2%)
3b	29 (13.5%)	28 (10.8%)
4	0	1 (0.4%)
Gleason score		
2-6	106 (49.3%)	75 (28.8%)
7	81 (37.7%)	119 (45.8%)
8-10	28 (13.0%)	66 (25.4%)
Median prehormone PS concentration,	11.3 (0.4-57.0) A	15.0 (1.8-59.6)
Indeed "Median initial PSA concentration" ug/L (range)	SIMILA	R GROUPS

Treatment

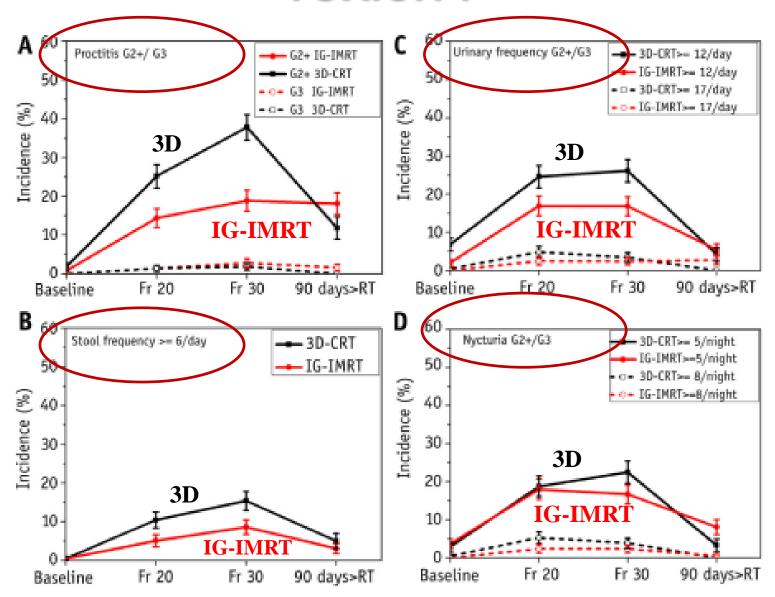
Risk category	3D	IG-IMRT
Low	34 (15.8%)	
Intermediate	72 (33.5%)	75 (28.8%)
High	109 (50.7%)	185 (71.2%)
Seminal vesicle dose	(Gy)	
0	43 (20.0%)	50 (19.2%)
50	35 (16.3%)	0
68	101 (47.0%)	0
70	0	125 (48.1%)
78	36 (16.7%)	85 (32.7%)
Planning margins (m	m)	
5	0	107 (41.3%)
6-8	0	151 (58.3%)
10	215 (100%)	1 (0.4%)
Hormone therapy	42 (19.5%)	174 (66.9%)
TURP	24 (11.2%)	28 (10.8%)
Diabetes mellitus	12 (5.6%)	29 (11.2%)
Abdominal surgery	57 (26.5%)	65 (25.0%)
Smoking	34 (15.8%)	28 (15.0%)



Mean dose and 10th to 90th percentiles are shown



TOXICITY





Prostate IGRT

ARTICLE IN PRESS

Radiotherapy and Oncology xxx (2014) xxx-xxx



Contents lists available at ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



Original article

Is "pelvic radiation disease" always the cause of bowel symptoms following prostate cancer intensity-modulated radiotherapy?

Myo Min^{a,*}, Benjamin Chua^a, Yvonne Guttner^d, Ned Abraham^d, Noel J. Aherne^a, Matthew Hoffmann^b, Michael J. McKay^{c,*,1}, Thomas P. Shakespeare^{a,1}

*North Coast Cancer Institute, Coffs Harbour; b North Coast Cancer Institute, Port Macquarie; c North Coast Cancer Institute, Lismore; d Coffs Harbour Hospital, Australia



Prostate IGRT

- Multicenter study
- N=102
- Prostate cancer
- Bowel symptoms persisting >90 days post-RT
- IMRT-IGRT
- Dose: 74-78 Gy at 1,8-2 Gy/fx
- Bowel symptoms + ENDOSCOPIC EXAMINATION
- Endoscopy findings:

56% Polyps

49% Diverticular disease

38% Haemorrhoids

29% radiation proctopathy with associated pathology

4% radiation proctopathy alone

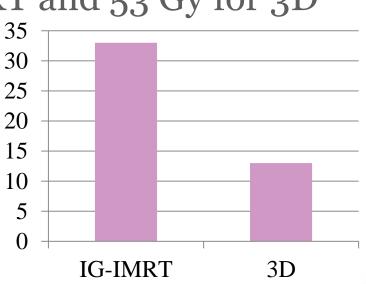


Clinical trial, Phase II (hepatocellular carcinoma)

- IG-IMRT (N=65) vs 3D (N=122)
- Stage III-IV
- Period: 2006-2011
- Retrospective
- Dose: 62 Gy for IG-IMRT and 53 Gy for 3D



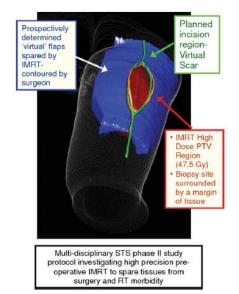
Survival at 3 years:

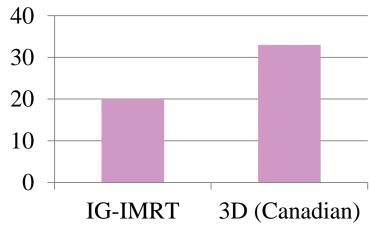




Clinical trial, Phase II (Sarcoma)

- Single Institution study
- Lower extremity soft tissue sarcoma
- N=56
- Period: 2005-2009
- IG-IMRT
- Dose: 50 Gy at 2 Gy/fx
- Acute wound complication:
- Local control 88%
- OS: 74%







Clinical trial, Phase II (Lung)

- Multicenter study
- Prospective
- Inoperable T1/T2 NSCLC
- N=60
- Period: 2003-2005
- SBRT
- Dose: 45 Gy at 15 Gy/fx
- Grade 3 toxicity: 21%
- Local control 96%
- OS: 65%

Table 3							
Lung-related	toxicity	maximum	grade	per	patient	number	of
affected pati	ents						

Toxicity	CVD (17 patients)		COPD (40 patients)	
	Gr 1-2	Gr 3	Gr 1-2	Gr 3
Cough	4	-	11	1
Dyspnoea	2	2	8	2
Pneumonia	_	_	1	1
Pneumonitis	3	_	7	_
Fibrosis	8	1	12	1
Atelectasis	3	1	3	_
Pleural effusion	6	2	5	-
Heart disorder	1	-	-	1
Esophagitis	1	-	1	-

Toxicity grading was done according to CTC v2. Radiation-related pulmonary fibrosis >90 days post-treatment was graded according to RTOG/EORTC Late Radiation Morbidity Scoring Scheme.

Maximum grade refers to the highest degree of toxicity recorded during follow-up. CVD, cardiovascular disease; COPD, chronic obstructive pulmonary disease.



Clinical trial, Phase II (oligometastases)

- Single Institution study
- Oligometastases
- N=25
- Period: 2004-2006
- SBRT
- Dose: 50 Gy at 5 Gy/fx + sunitinib
- Grade 3 toxicity: 28%
- Local control 75%
- OS: 71%; PFS: 56%

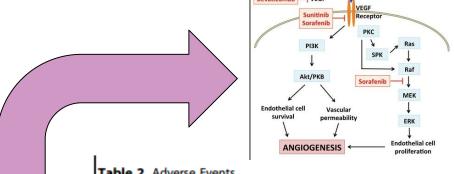


Table 2. Adverse Events.

Adverse Event	All grades	Grade 3	Grade 4	Grade 5
Anemia	18	2	0	0
Neutropenia	14	2	0	0
Fatigue	18	0	0	0
LFT abnormalities	15	1	0	0
Thrombocytopenia	15	4	0	0
Mucositis/stomatitis	8	0	0	0
Nausea/vomiting	7	0	0	0
Skin changes	4	0	0	0
Diarrhea	5	0	0	0
Hypertension	3	0	0	0
Bleeding	4	1	0	1*
Metabolic abnormalities	2	1 (PO ₄)	0	0
Increased creatinine	5	0	0	0

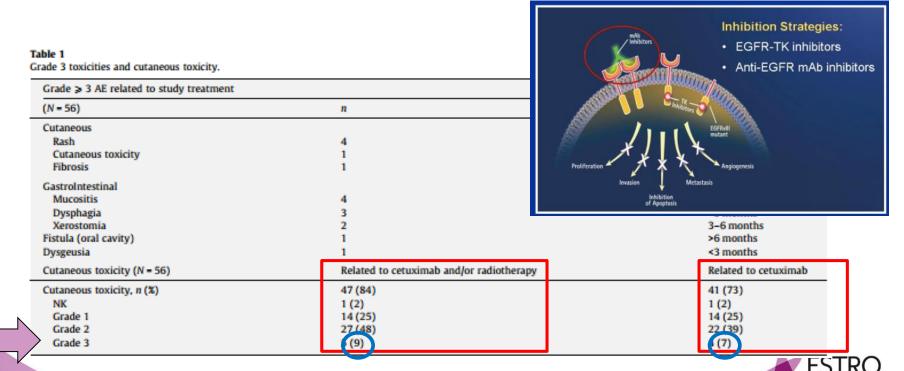
*One case occurred after sunitinib treatment and was likely related to reirradiation performed prior to protocol therapy. doi:10.1371/journal.pone.0036979.t002

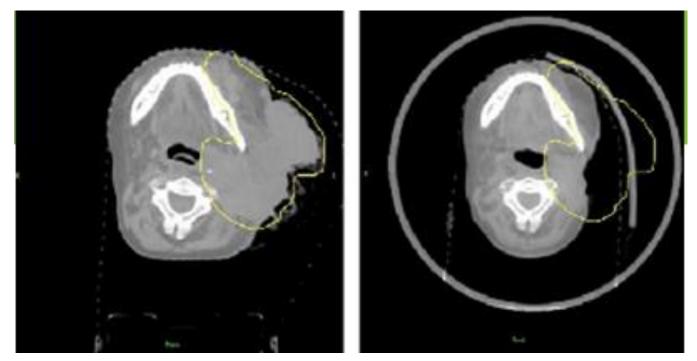


Clinical trial, Phase II (head and neck)

- Multicenter study
- Reirradiation
- N=60
- Period: 2007-2010
- SBRT

- Dose: 36 Gy at 6 Gy/fx + cetuximab
- Grade 3 toxicity: 18%
- OS: 47,5%





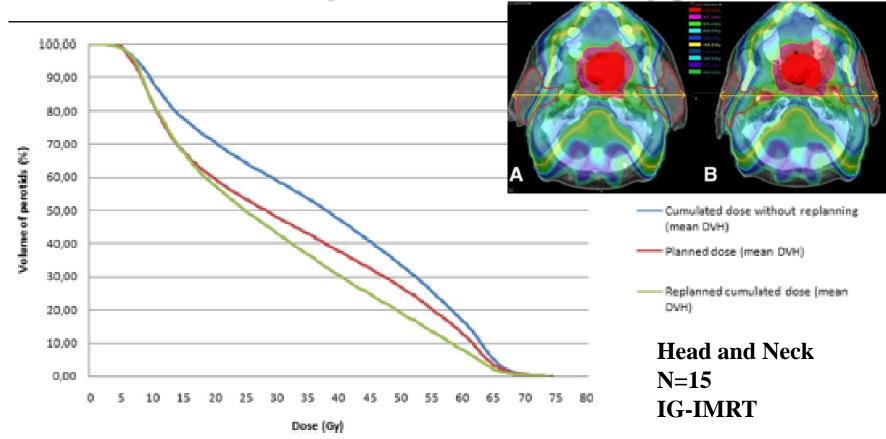
Corresponding axial CT slices from the beginning and the end of treatment.

The volume of the PTV changed from 606 to 336 cm3 over treatment, a decrease of 45%.

Spinal cord Do5 differed from the planned value by: 3.5% (average) +/- 9.8% (standard desviation)



Adaptive radiotherapy



Replanning decreased the PG mean dose by 5 Gy, and 11% the xerostomia

LEVEL 2B: INDIVIDUAL COHORT STUDY

ESTRO School

FUTURE DIRECTIONS (phase III studies on-going)

- A Randomised, Two Centre Trial on Daily Cone-beam vs Standard Weekly Orthogonal IGRT for **Prostate**
- Hypofractionated IGRT in Patients With Stage II-III Non-Small Cell Lung Cancer
- Biological Image Guided Antialgic SBRT of Bone Metastases: a Randomized Phase II/III Trial
- Evaluation of 3DCRT Versus IGRT and Analysis of Early Response in Head and Neck Cancer.
- Tomotherapy vs Conventional Radiation for Adjuvant Pelvic RT in Ca Cervix.
- Can 3D Ultrasound Be Used Reproducibly by RTTs in Partial Breast IGRT?



BEYOND IGRT:

Daily real time planning (RTP)— Treatment of prostate cancer, clinical implementation, and technique

- 60 RTP's were delivered (10 daily RTP/patient) in 6 consecutive patients.
- In 20% of the cases, the CTV-DVH by RTP improved by >10%.

Plans V40 (%) V50 (%) V60 (%) V70 (%) RTP 47.1 29.9 18.6 7.8

IGRT 63.8 49.3 38.0 26.5

20%



FUTURE DIRECTIONS

Radiotherapy and Oncology 109 (2013) 165-169



Contents lists available at SciVerse ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



Learning methods in radiation oncology

The utility of e-Learning to support training for a multicentre bladder online adaptive radiotherapy trial (TROG 10.01-BOLART)



Farshad Foroudi ^{a,*}, Daniel Pham ^a, Mathias Bressel ^b, David Tongs ^a, Aldo Rolfo ^{a,1}, Colin Styles ^a, Suki Gill ^a, Tomas Kron ^a

^a Division of Radiation Oncology and Cancer Imaging, Peter MacCallum Cancer Centre, Melbourne, Australia; ^b Centre for Biostatistics and Clinical Trials, Peter MacCallum Cancer Centre, Melbourne, Australia



IGRT confidence and knowledge

- To demonstrate the utility of an e-Learning programme for providing training regarding a multi-centre **IGRT** trial.
- Participants : **185 RTTs** from 12 centres.
- There was **an increase confidence after** modules (p < 0.001).
- The pre scores increased from $67 \pm 11 \implies 79 \pm 8 \text{ (p < 0.001)}$



IGRT confidence and knowledge

Confidence questions	Pre e-Learni	ng	Post e-Learni	ng	<i>p</i> -Value
	n	Percentage	n	Percentage	
Identifying bladder on CT					< 0.001
Not confident	0	0.0	0	0.0	
A little confident	20	10.8	4	2.2	
Somewhat confident	60	^{32,4} 57%	31	31 % 16.8	
Quite confident	69	37.3	102	55.1	
Very confident	36	19.5	48	25.9	
Identifying soft tissue anatomies of	n pelvic CBCT				< 0.001
Not confident	13	7.0	0	0	
A little confident	49	26.5	7 💂	3.8	
Somewhat confident	80	25.5	0 71 5	60_0 38.4	
Quite confident	39	21.1	91	49.2	
Very confident	4	2.2	16	8.6	
Implementing the BOLART at your	centre				< 0.001
Not confident	28	15.1	1	0.5	
A little confident	40	21.6	9	4.9	
Somewhat confident	58	314 32%	48 69	0/0 25.9	
Quite confident	45	24.3	86	46.5	
Very confident	14	7.6	41	22.2	

E-LEARNING WAS FEASIBLE AND IMPROVED CONFIDENCE AND KNOWLEDGE



CONCLUSIONS: Why IGRT?

- Security
- Precision
- Accuracy (dose escalation)
- Homogeneity
- Potentially, less toxicity: clinical trials needed!!
- Reliability
- Adapt to changes in antomy
- Shortening RT





Case report: Cervix



Sofia Rivera, Gustave Roussy, Villejuif, France

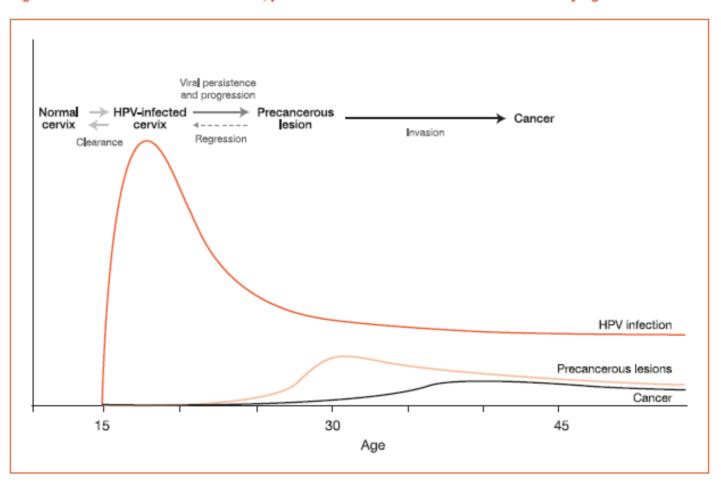


Case from the Gyn GEC ESTRO Network / FALCON WS Courtesy of Pr Pötter

Advanced skills in modern radiotherapy
June 2017

HPV infection natural history

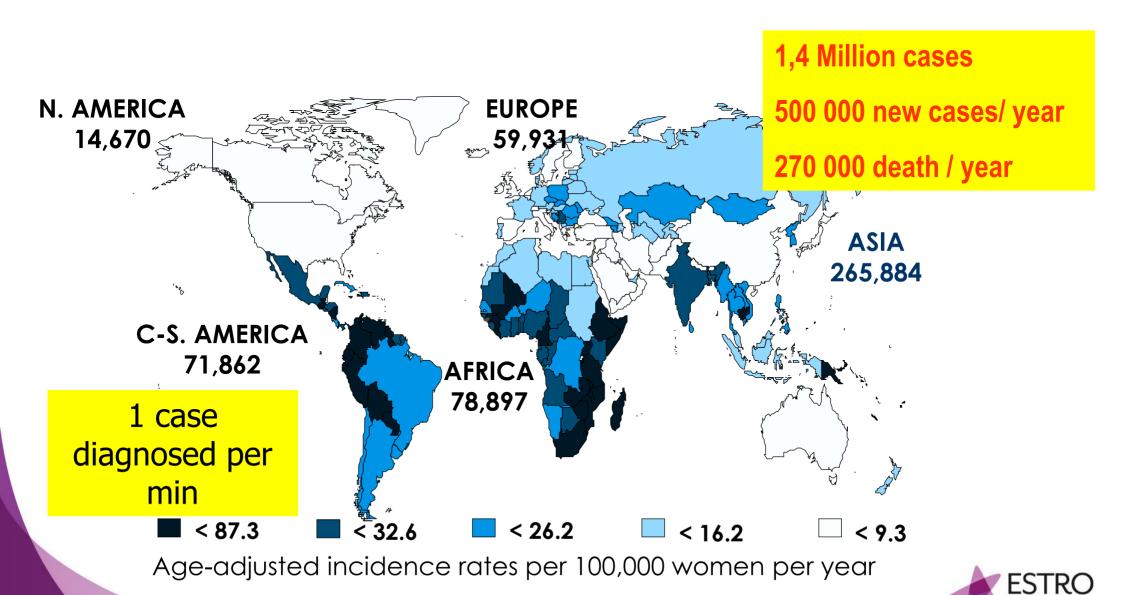
Figure 1. Prevalence of HPV infection, precancerous lesions and cervical cancer by age of women



Source: Schiffman M, Castle PE. The promise of global cervical-cancer prevention. New England Journal of Medicine, 2005, 353(20): 2101–2103. (© 2005 Massachusetts Medical Society. Adapted with permission.)



Cervix cancer diagnosis OMS (2002)



Patient History

- •42-year old woman.
- •WHO performance status=0
- No clinical symptom
- •No palpable node
- •Squamous cell carcinoma, grade 3
- •TNM: T3b N1 M0



Clinical findings of gyn. examination: at DIAGNOSIS Infiltrative Exophytic Cervix Vagina Parametria **Dimensions:** Width: 90 mm Thickness: 50 mm Height: 60 mm Vaginal inv.: 20 mm **ESTRO** Gyn contouring workshop Barcelona, May 2012 *Radiological findings integrated on the drawing



Clinical findings of gyn. examination: SUMMARY

FIGO stage: IIIB

	At diagnosis	At brachytherapy
Width	90 mm	
Thickness	50 mm	
Height*	60 mm	
Left parametrium	Infiltration to pelvic wall	
Right parametrium	Proximal infiltration	
Vagina	20 mm: left & posterior wall	
Bladder**	Not infiltrated	
Rectum**	Not infiltrated	

ESTRO

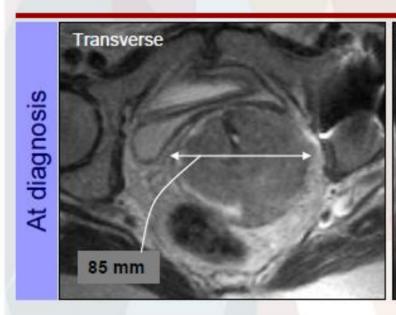
Gyn contouring workshop Barcelona, May 2012

*Some uncertainty in assessment of height

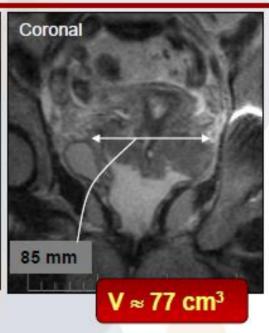
**Endoscopy at diagnosis



MRI findings







Comment:

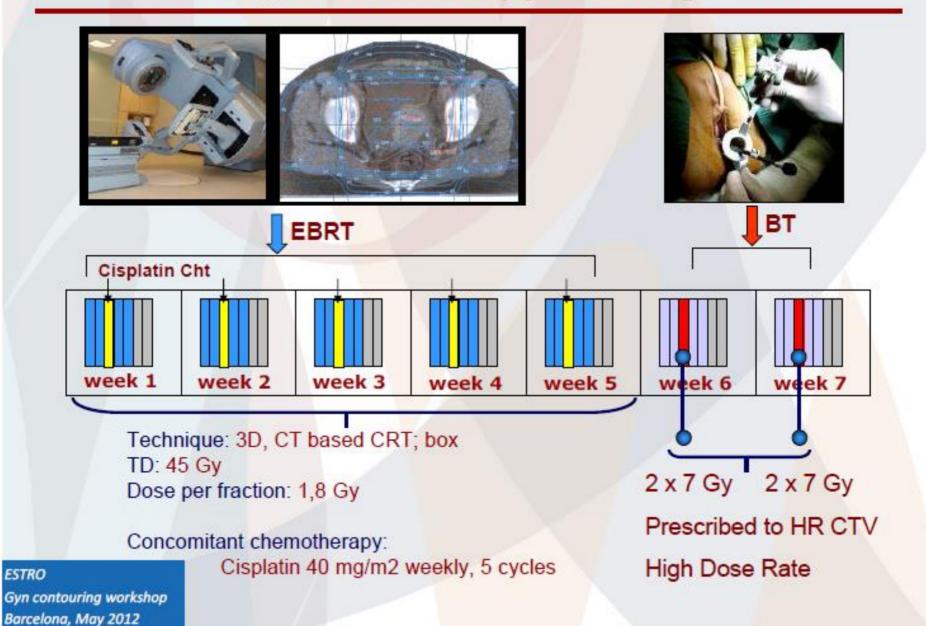
Only the representative slices are shown here. Use the information from complete initial MRI data set to fully understand the extent and topography of the tumour.

ESTRO

Gyn contouring workshop Barcelona, May 2012



EBRT, Chemotherapy & timing of BT





Clinical findings of gyn. examination: at Brachytherapy Infiltrative Exophytic Cervix Vagina



Dimensions:

Width: 70 m

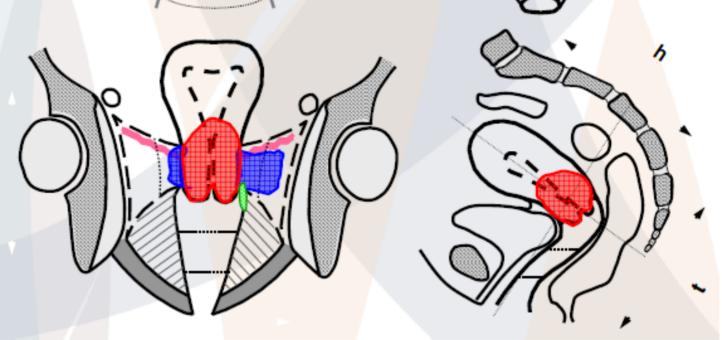
Thickness: 40

Height: 40

Vaginal inv.: 10

ESTRO

Gyn contouring workshop Barcelona, May 2012



necrosis



Clinical findings of gyn. examination: SUMMARY

FIGO stage: IIIB

	At diagnosis	At brachytherapy
Width	90 mm	70 mm
Thickness	50 mm	40 mm
Height*	60 mm	40 mm
Left parametrium	Infiltration to pelvic wall	Distal infiltration (≈ 30 mm)
Right parametrium	Proximal infiltration	Proximal infiltration (≈ 10 mm)
Vagina	20 mm: left & posterior wall	10 mm: left fornix
Bladder**	Not infiltrated	NA
Rectum**	Not infiltrated	NA

ESTRO

Gyn contouring workshop Barcelona, May 2012

*Some uncertainty in assessment of height

**Endoscopy at diagnosis



Brachytherapy application



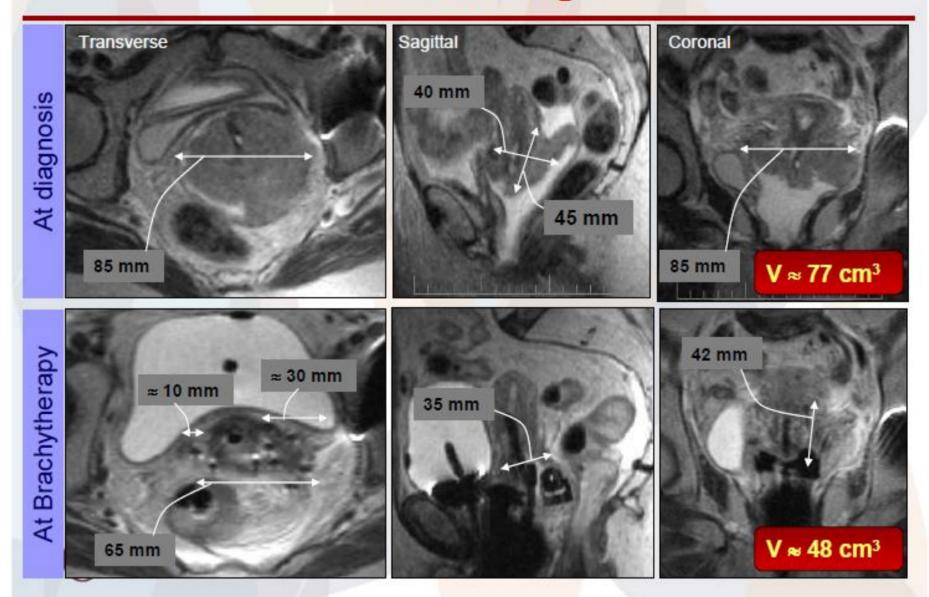


Tandem & Ring
Interstitial parametrial needles according to tumour spread

ESTRO Gyn contouring workshop Barcelona, May 2012 → Following applicator insertion: pelvic MRI with the applicator in place



MRI findings





ESTRO project

Recommendations from gynaecological (GYN) GEC ESTRO working group (II): Concepts and terms in 3D image-based treatment planning in cervix cancer brachytherapy—3D dose volume parameters and aspects of 3D image-based anatomy, radiation physics, radiobiology

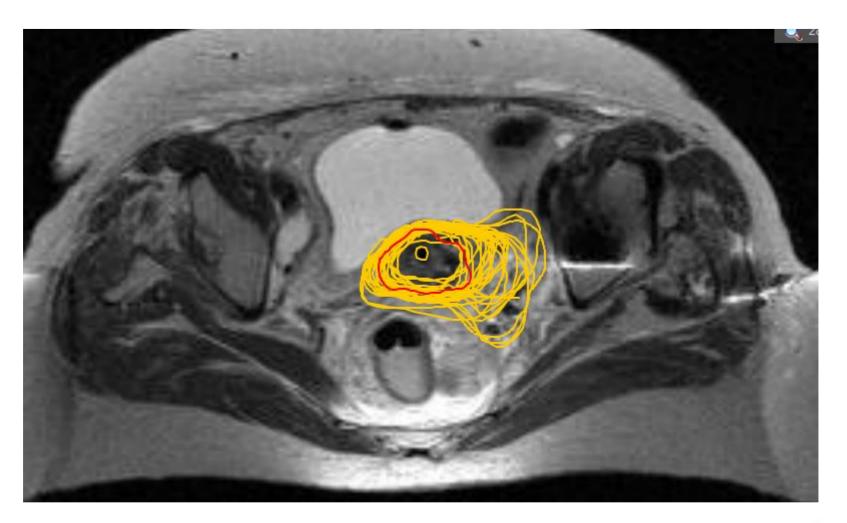
Richard Pötter^{a,*}, Christine Haie-Meder^b, Erik Van Limbergen^c, Isabelle Barillot^d, Marisol De Brabandere^c, Johannes Dimopoulos^a, Isabelle Dumas^b, Beth Erickson^e, Stefan Lang^a, An Nulens^c, Peter Petrow^f, Jason Rownd^e, Christian Kirisits^a

^aDepartment of Radiotherapy and Radiobiology, Medical University of Vienna, Austria, ^bDepartment of Radiotherapy, Brachytherapy Unit, Institut Gustave Roussy, Villejuif, France, ^cDepartment of Radiotherapy, University Hospital Gasthuisberg, Leuven, Belgium, ^dDepartment of Radiation Oncology, Centre George-Francois Leclerc, Dijon, France, ^eDepartment of Radiation Oncology, Medical College of Wisconsin, Milwaukee, WI, USA, ^fService de Radiodiagnostic, Institut Curie, Paris, France



Heterogeneity in contouring target volumes besides the use of guidelines

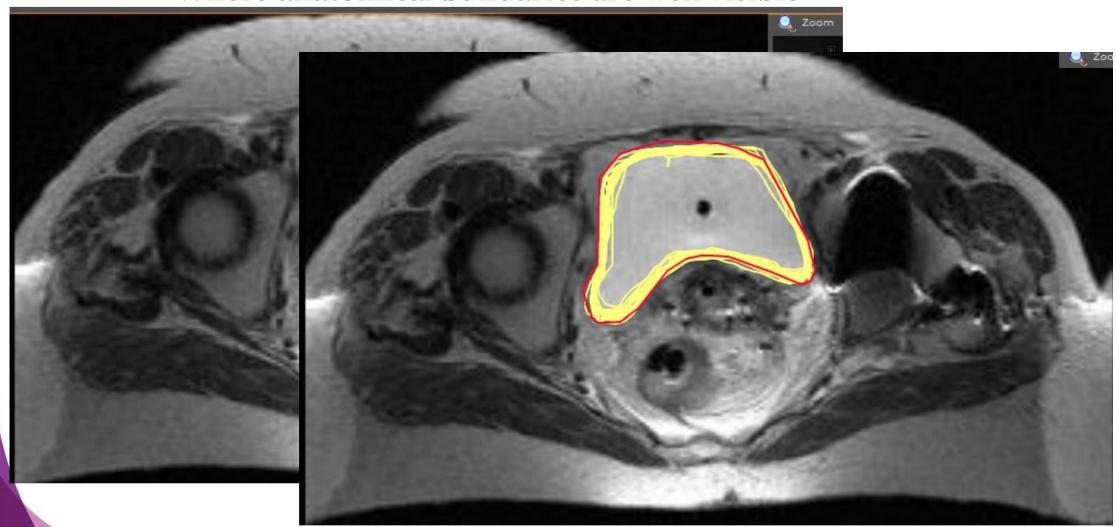
High Risk CTV





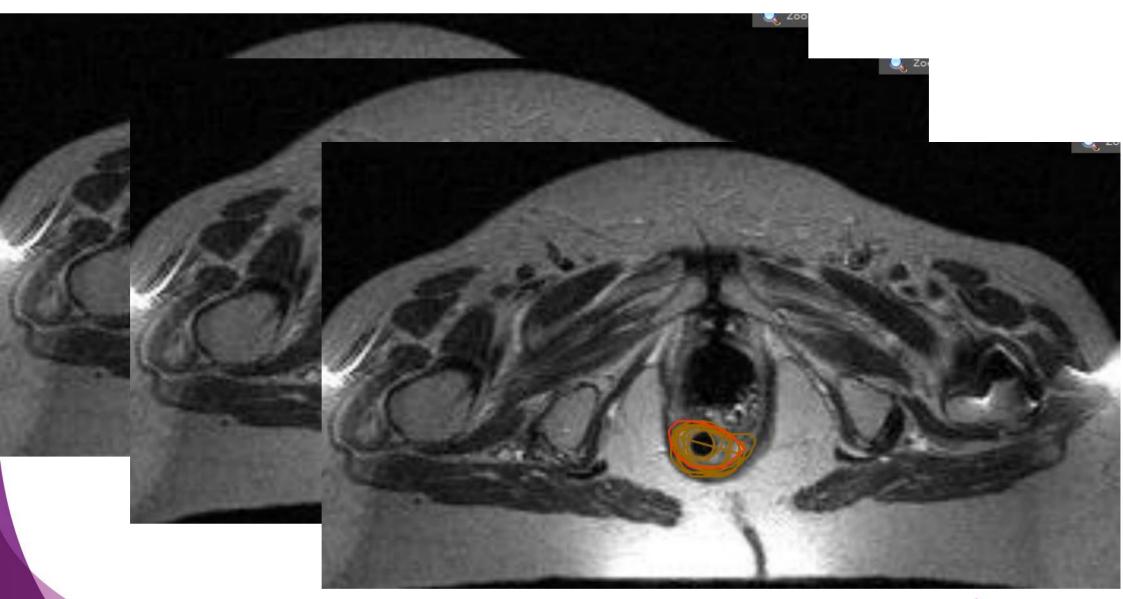
Quite good homogeneity in some OAR contouring

Where anatomical bundaries are well visible



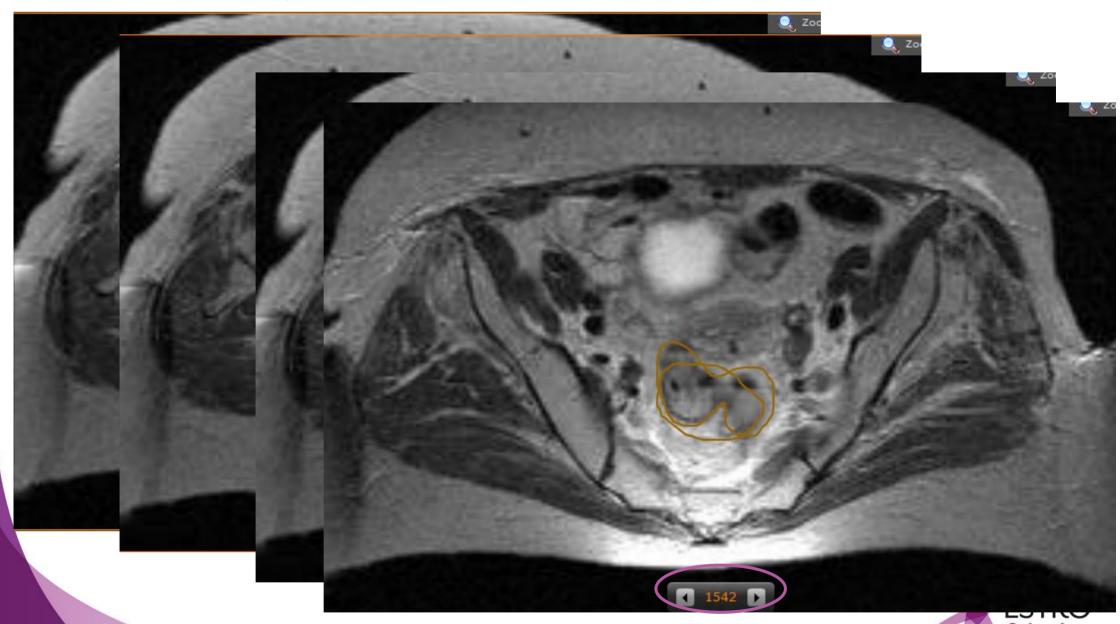


But it's not always the case!





Upper and lower limits are a source of heterogeneity in contouring as well



5 slices = 1,5cm difference in the upper limit of the rectum

Take home messages:

- High quality CT, MR imaging and clinical examination are crucial for contouring targets and OAR in the pelvic region
- High quality re-imaging and clinical examination are key points in cervical cancer to adapt contours for brachytherapy dosimetry
- MR is a key imaging modality in gynecology

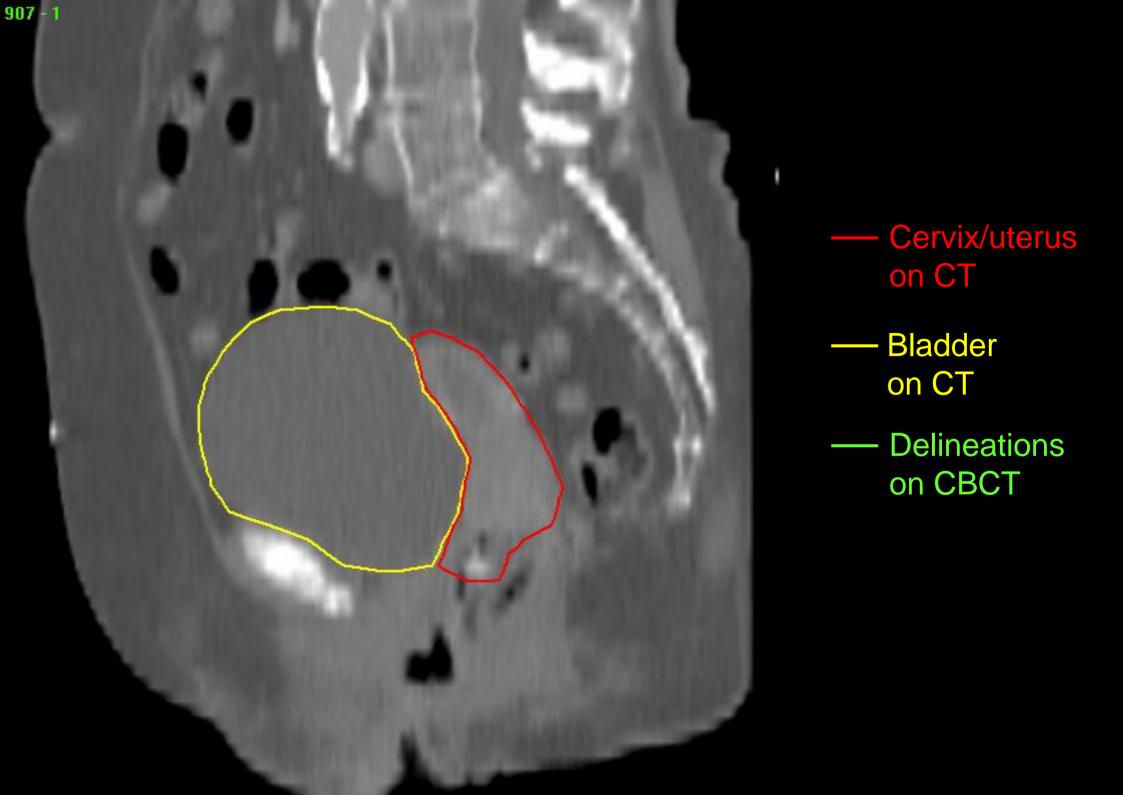




Cervix case – Physics aspects

Peter Remeijer





The main issue

- Very large movement
- Clinically a margin of 2 cm is used
- Probably too small



Options

- Drinking protocol
 - > Does it work?
- Scan and postpone treatment if not ok
 - Ask patient to empty bladder or drink and wait
 - Not very efficient
- Adapt the plan
 - Fast planning and delineation not readily available
 - ➤ A-priori plans select the right one at treatment



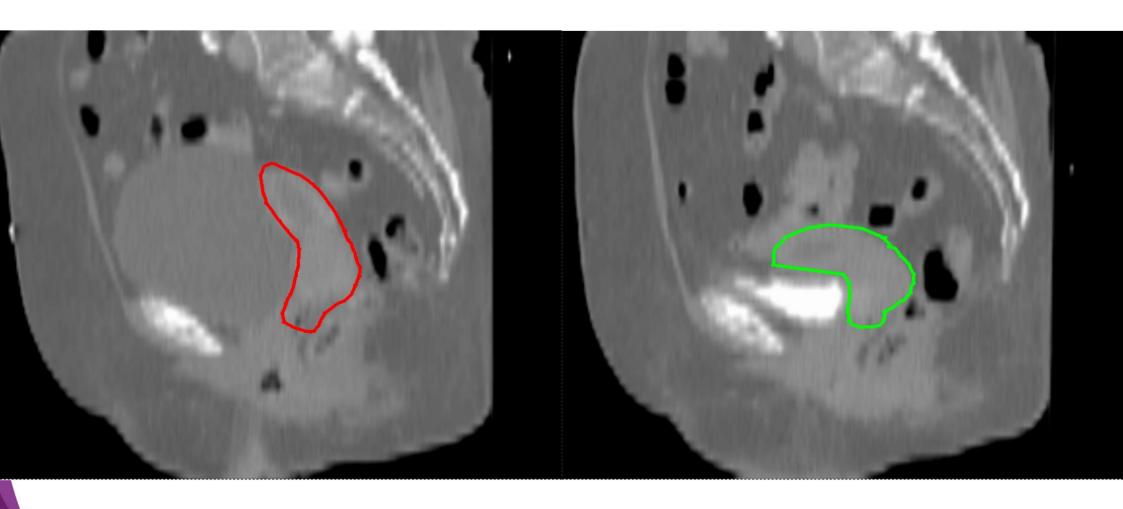


Options

- Drinking protocol
 - > Does it work?
- Scan and postpone treatment if not ok
 - Ask patient to empty bladder or drink and wait
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- Adapt the plan
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 - ➤ A-priori plans select the right one at treatment



Full/empty bladder CT





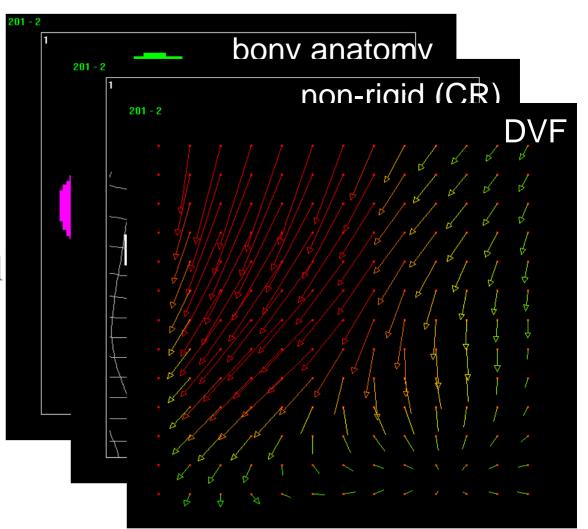
Generate uterus motion model



Delineate on full and empty bladder CT

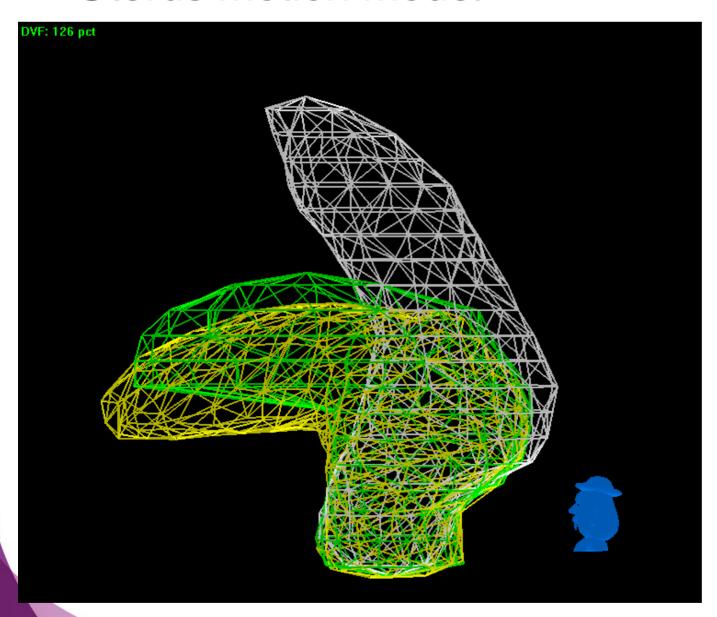
Deform full bladder contour

Generated warp field is model for organ motion





Uterus motion model

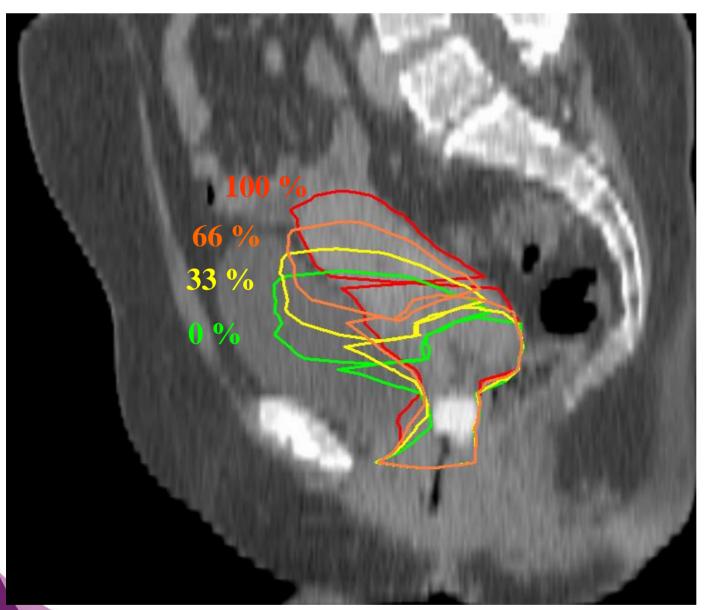


Select 4 bladder fillings based on this model:

- 0 %
- 33 %
- 66 %
- 100 %



Generated CTVs



Select 4 bladder fillings based on this model:

- 0 %
- 33 %
- 66 %
- 100 %



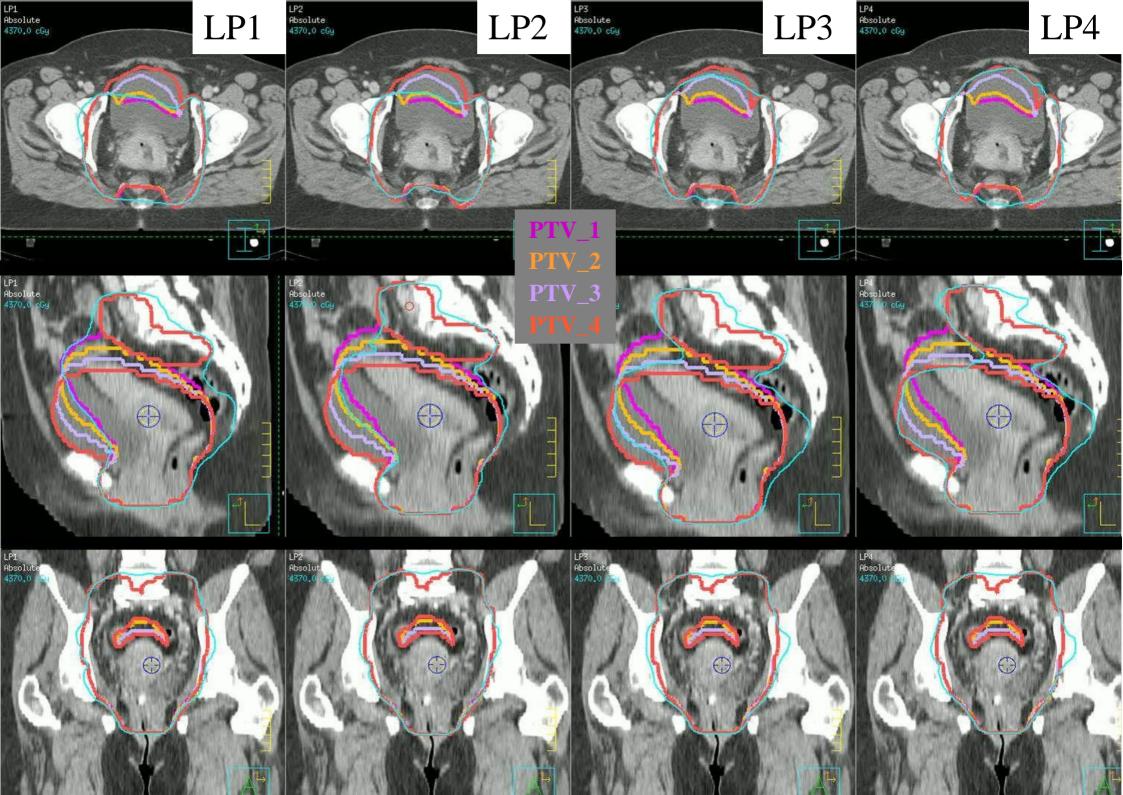
Planning



Planning

- 1 VMAT plan created manually (66%, LP2)
- Used as starting point to create three plans automatically (Pinnacle scripting)
- All plans based on full bladder CT scan
 - Dosimetrically not correct
 - Small deviations (1-2 %)

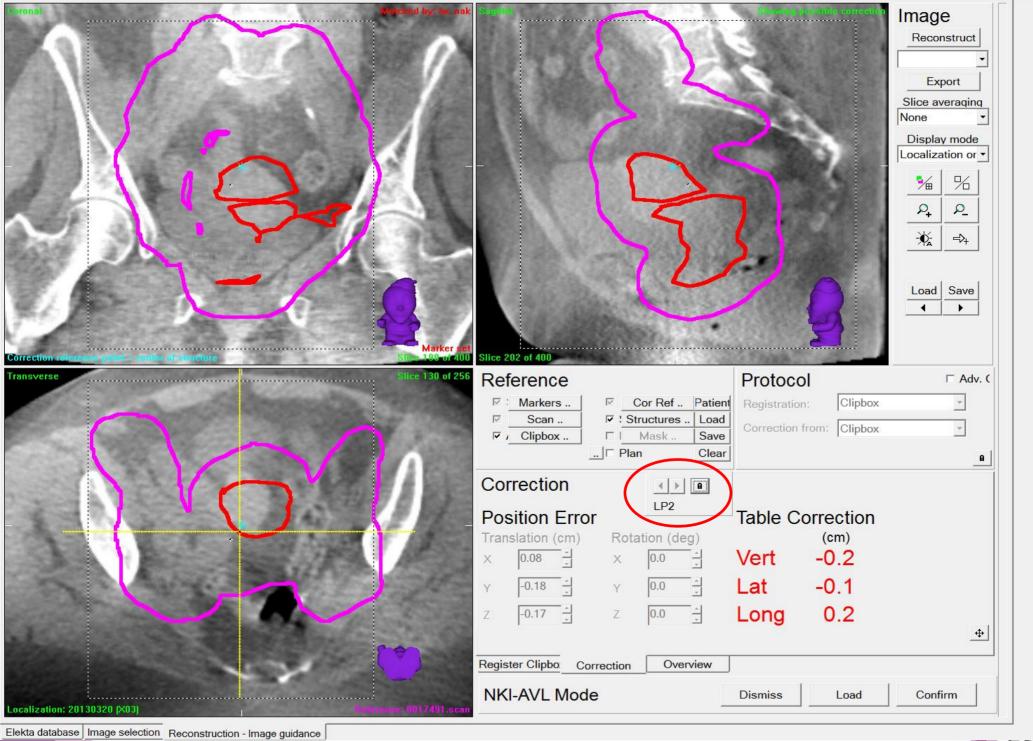




Treatment

- CBCT scan prior to treatment
- Select the 'best fitting structure'
- Select corresponding plan on the linac
- Treat
- All steps are checked/interlocked using in-house software





Alternatives?

- Only use full/empty scan
 - Margin about half the motion
 - Other errors small compared to motion



- > Import CBCT data in planning/delineation system
- > Delineate on scans, using planning scan as template
- Create plan library
- Protocolize everything. It is easy to make mistakes when doing planselection on the linac!





Take home messages

- Drinking protocols are usually not very effective
- Multiple scans to estimate the range of motion are
- Large improvements in geometrical accuracy can be achieved with simple methods
- Protocolize the plan selection process (error prone!)
- Planning and dosimetry is the least of your problems!





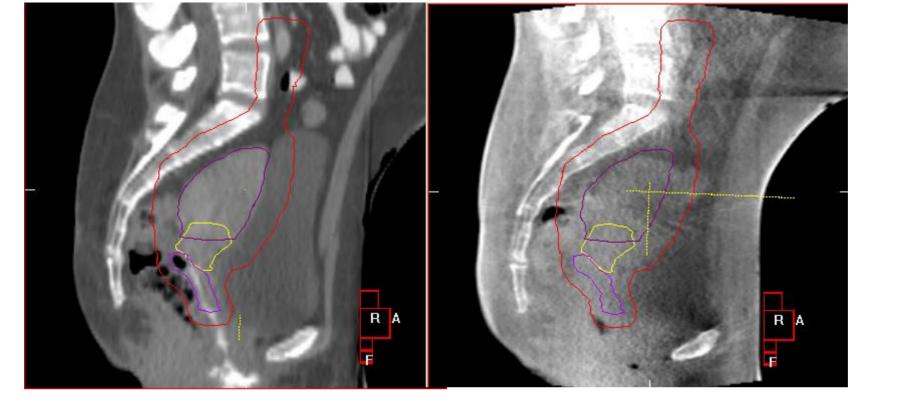


Cervix

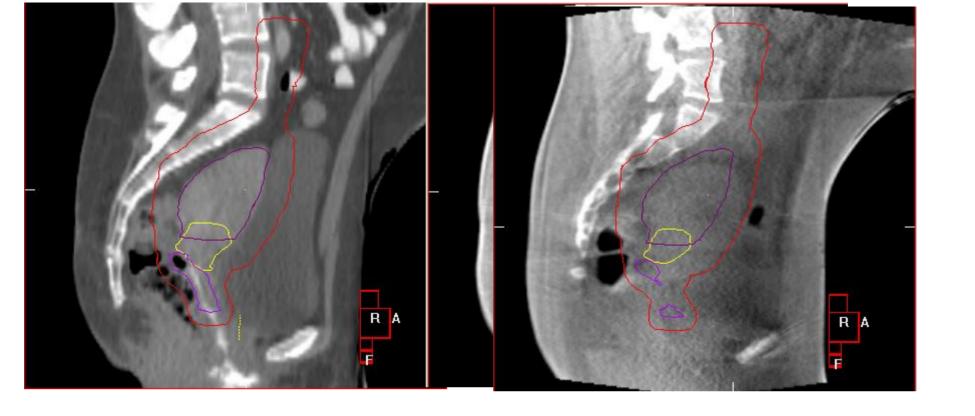
ART Library of Plans

Rianne de Jong *RTT*, Academic Medical Centre, Amsterdam Prague 2017

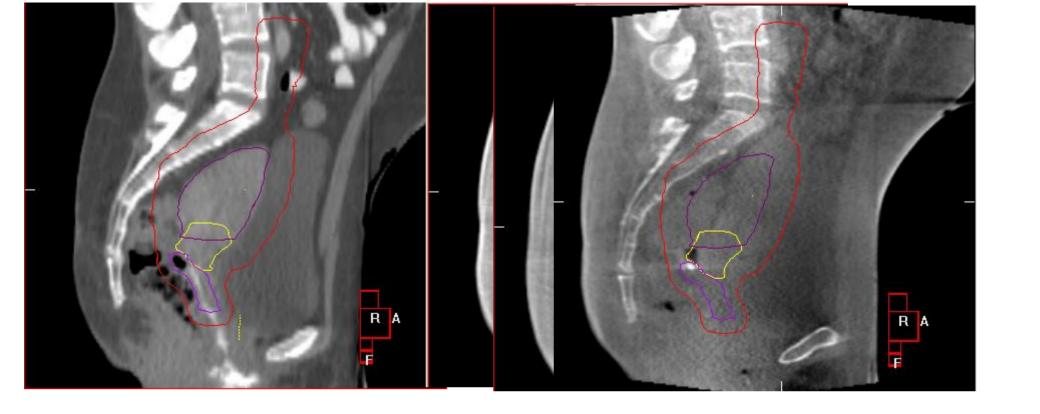




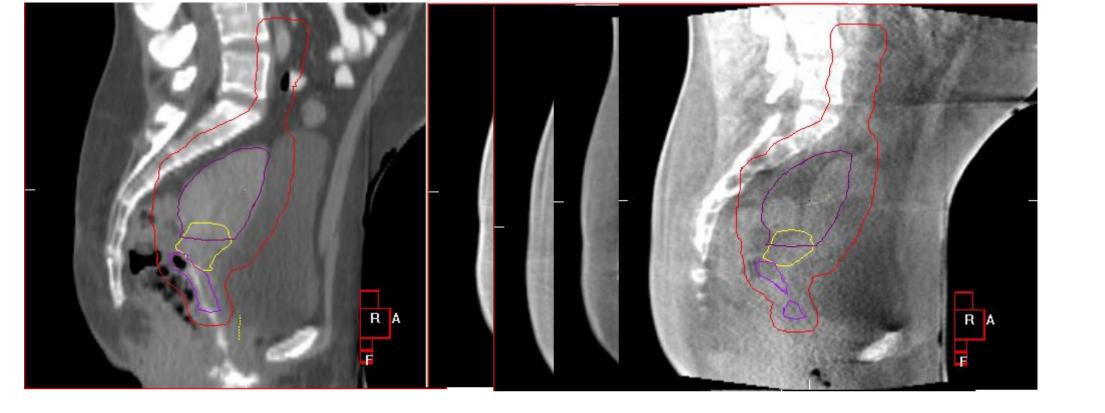




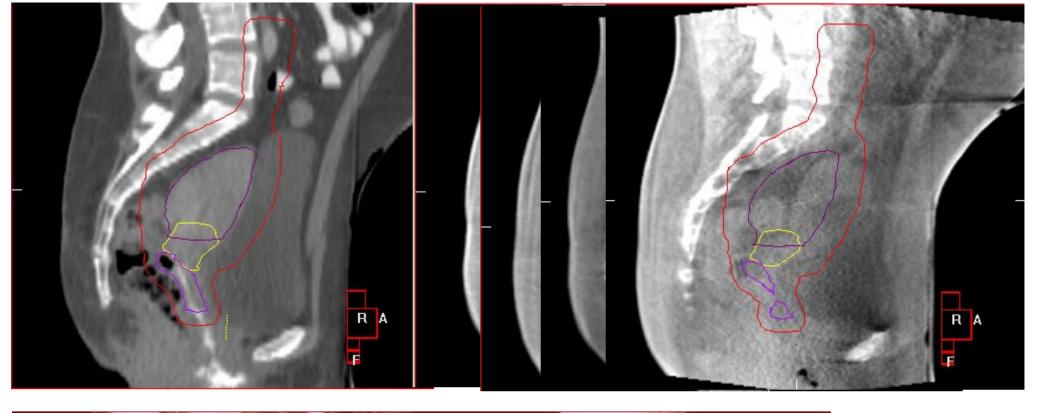


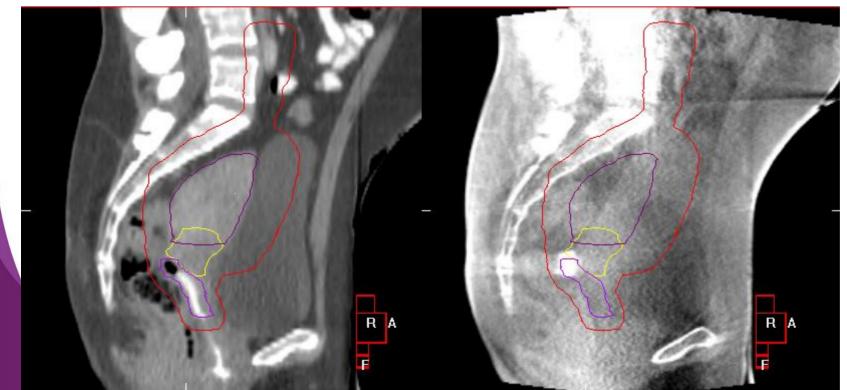




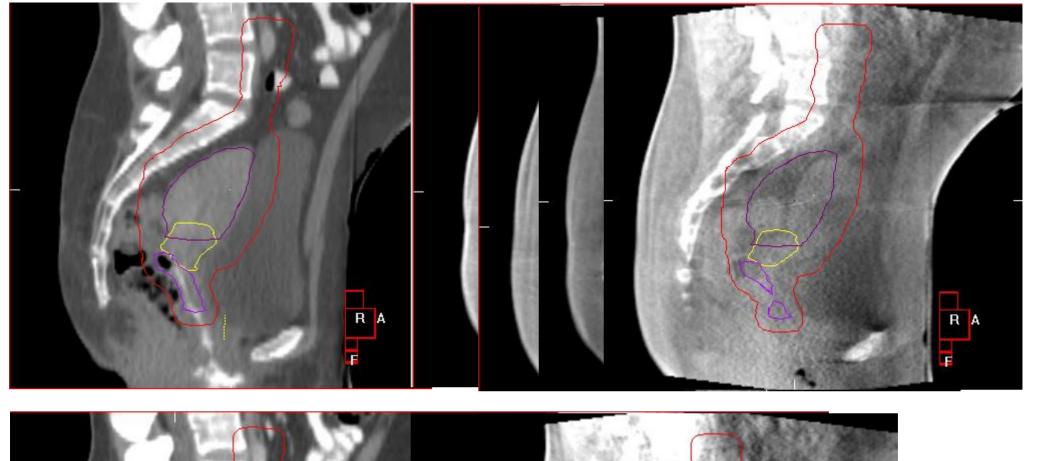






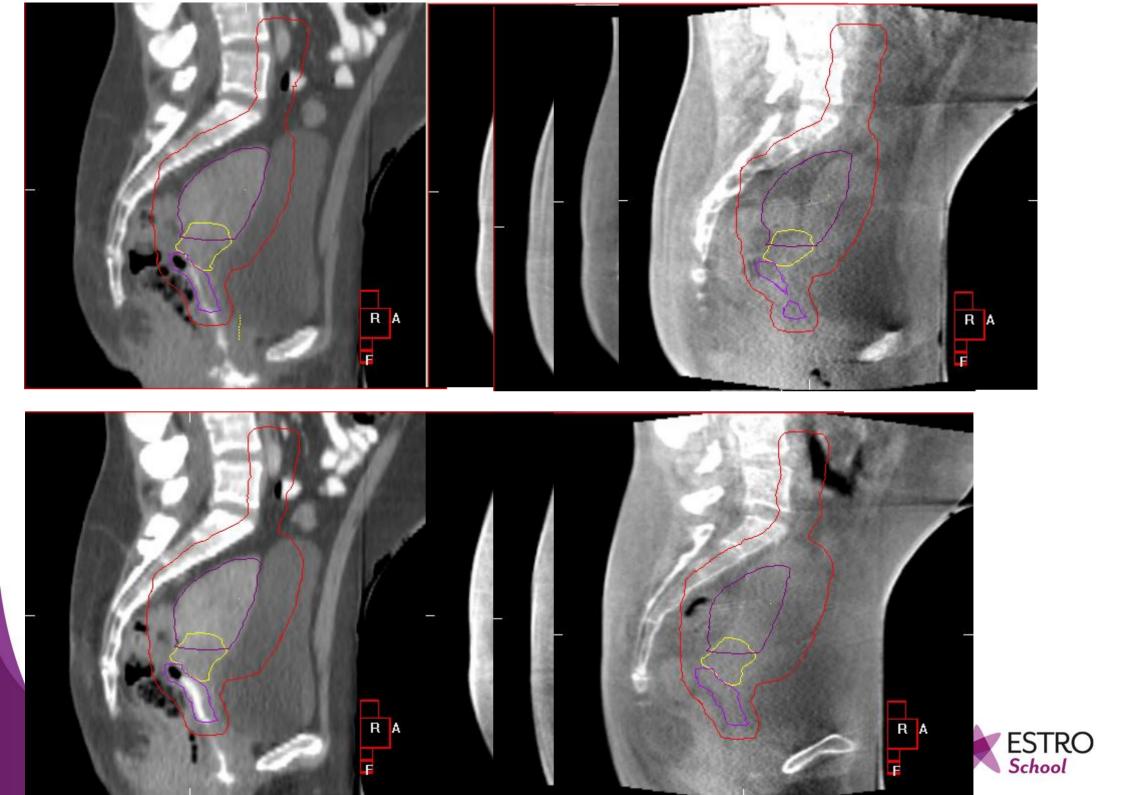


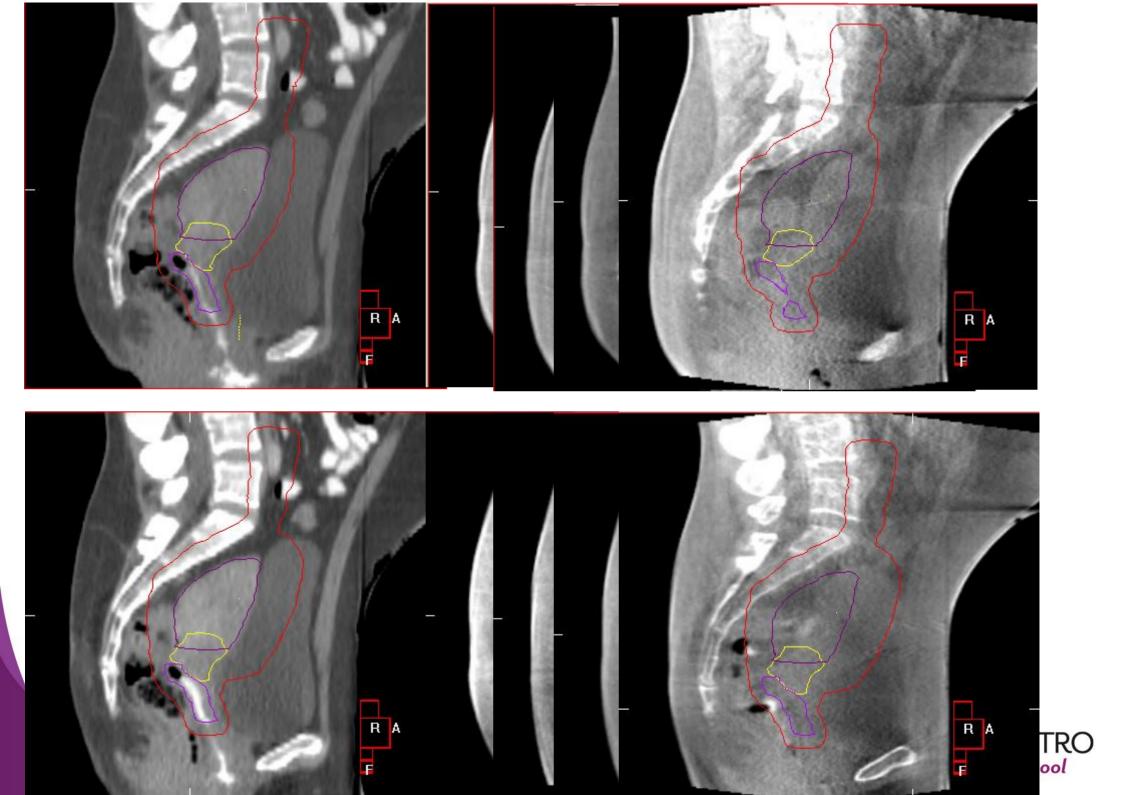


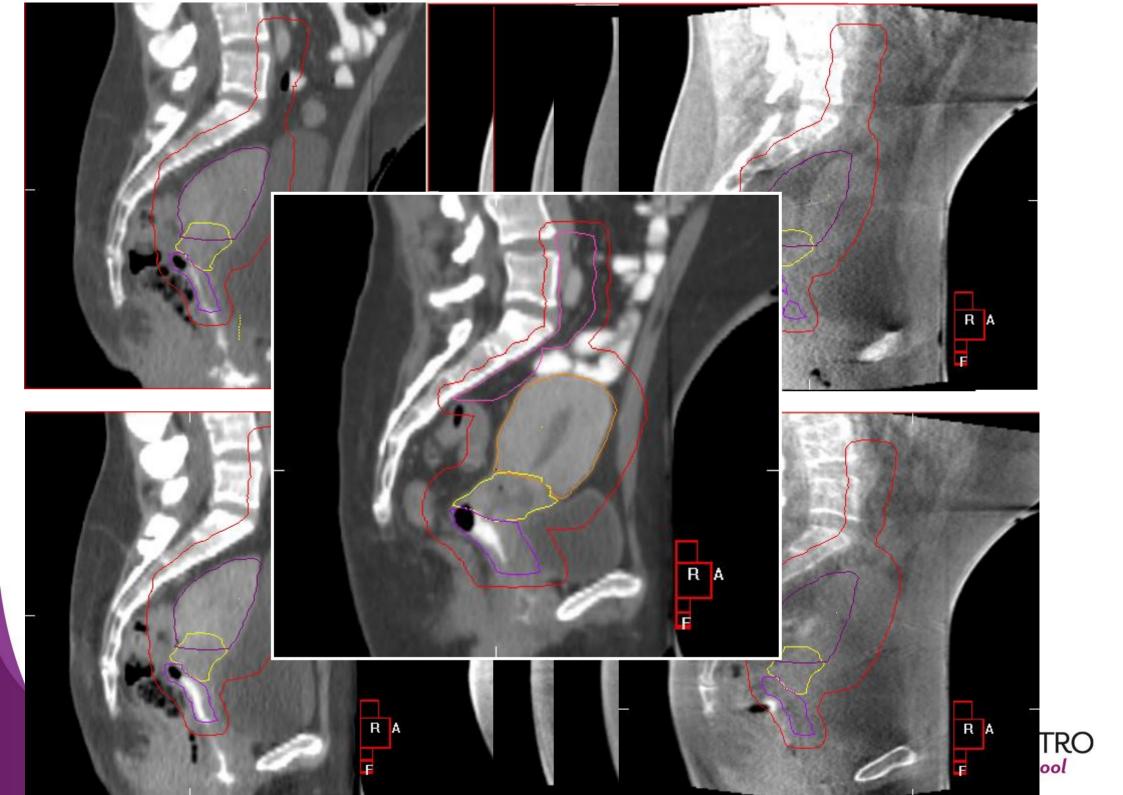


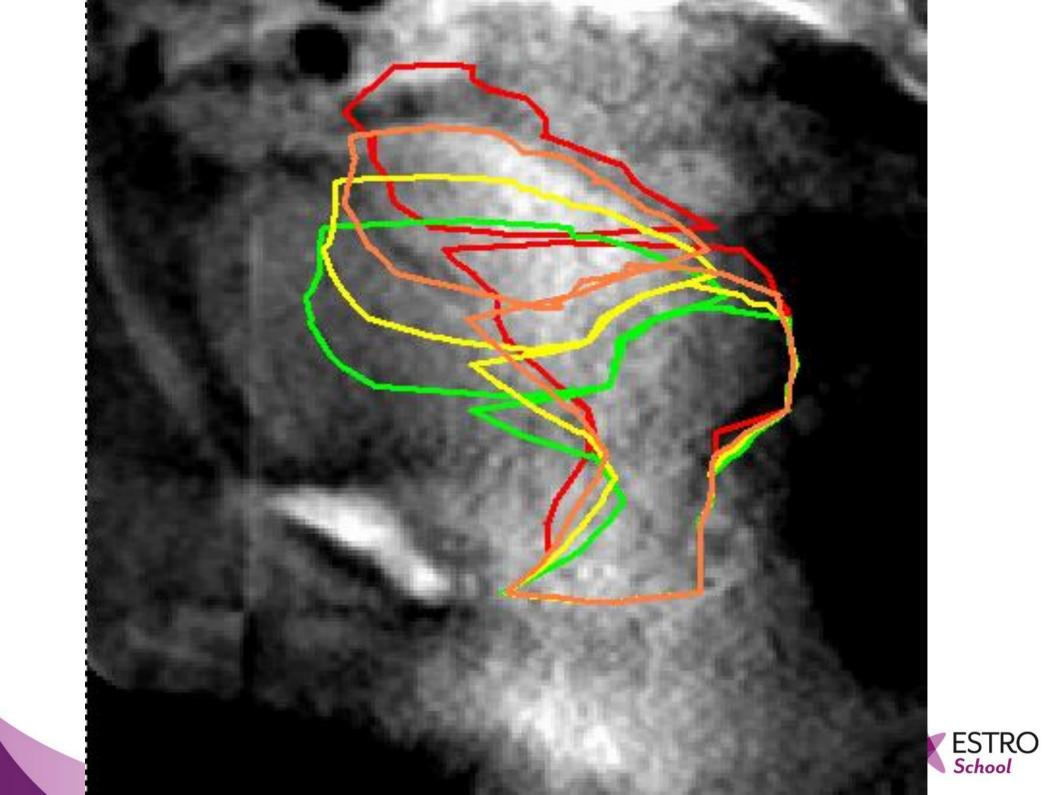


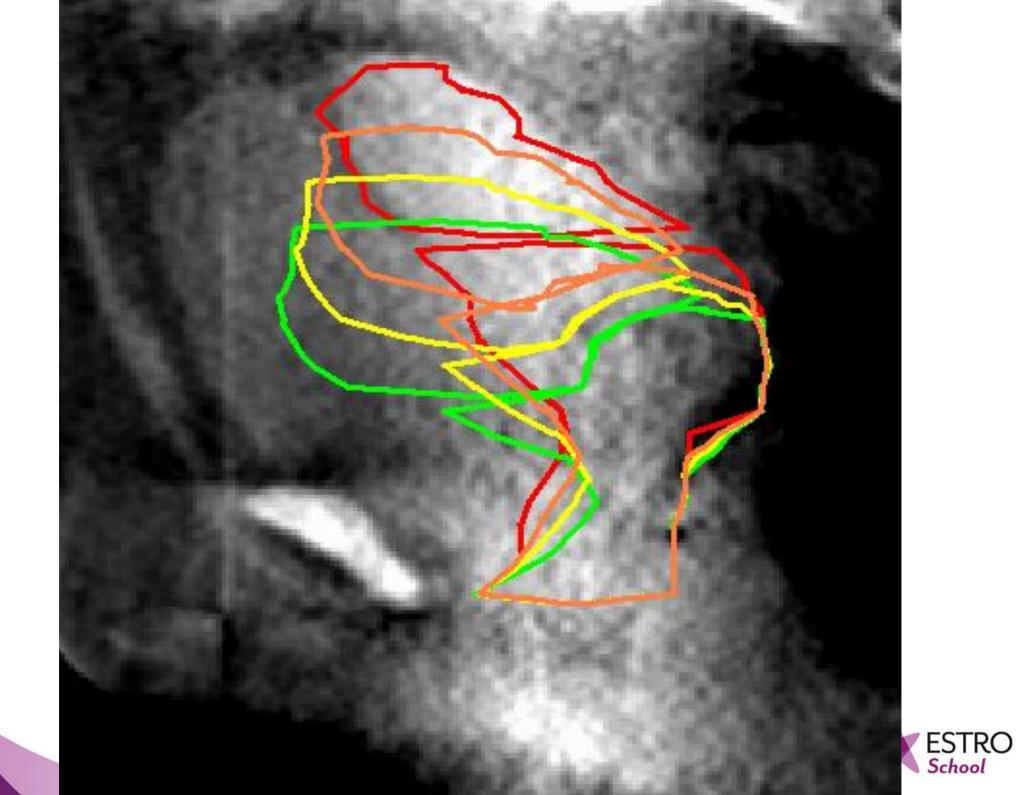


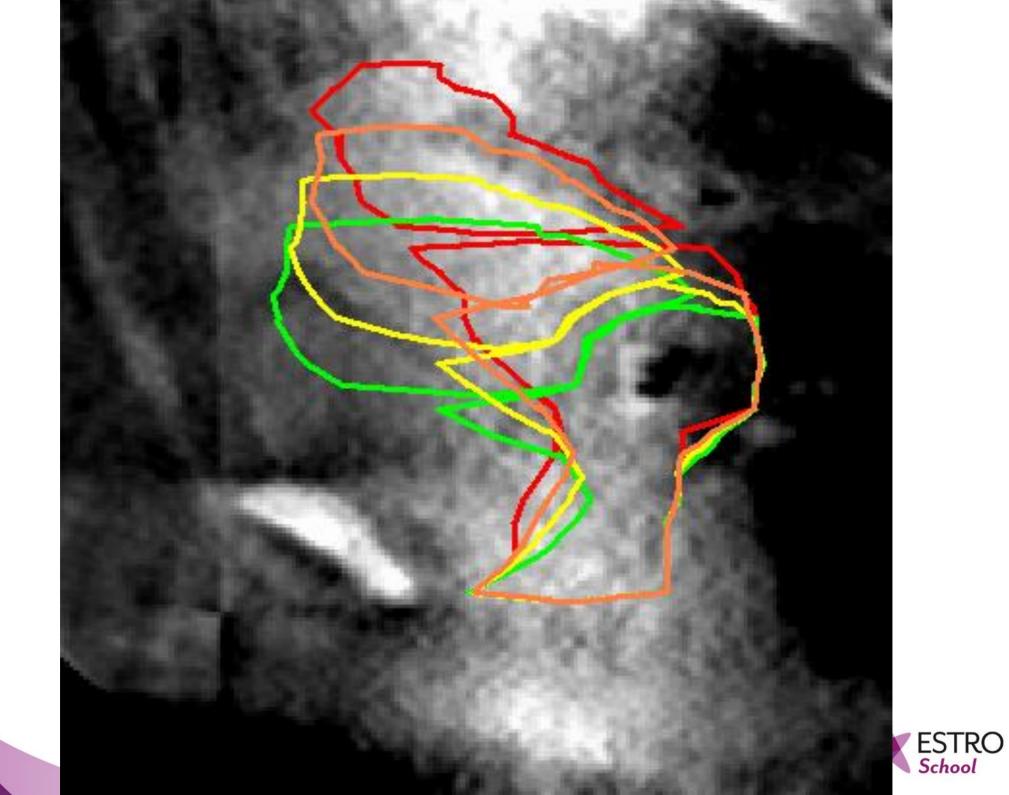












Question

Is it possible to consistently select a best fitting CTV based on CBCT scans?

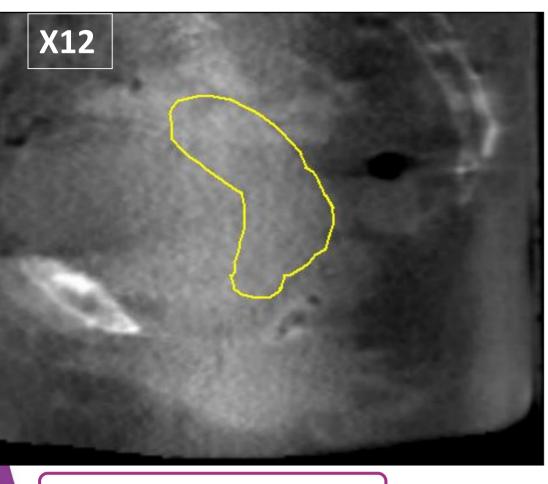


Observer study setup

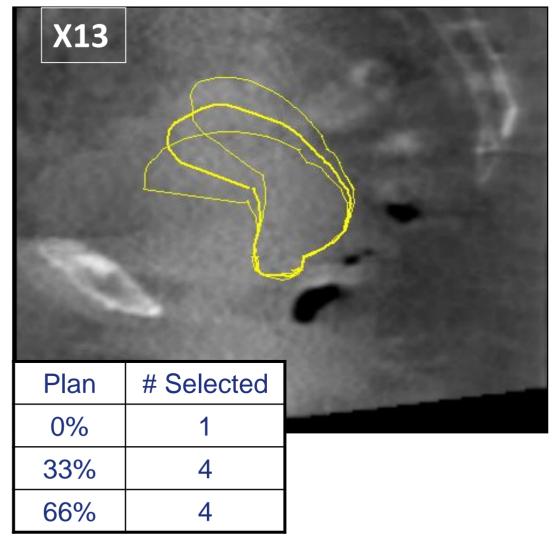
- 5 patients, 23 scans per patient (1 scan missing)
- Per patient 6 structures
 -20, 0, 33, 66, 100, 120%
- 9 observers (experienced RTTs)
- 2 sessions
- Workshop in between sessions to determine Gold
 Standard with observers: RTT, physicians and physicists



Patient 1



100% agreement

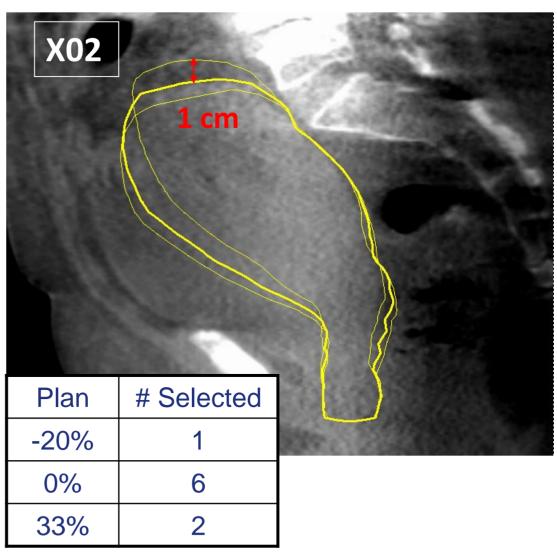




Patient 2

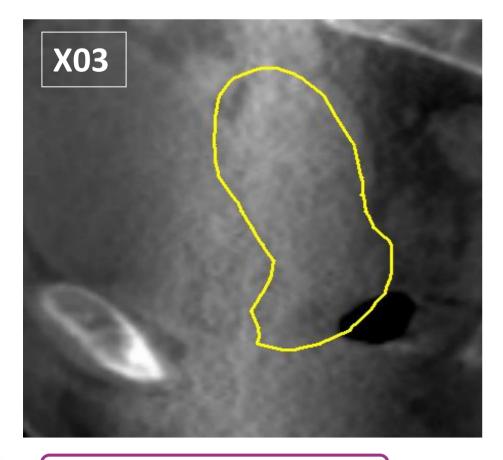


100% agreement

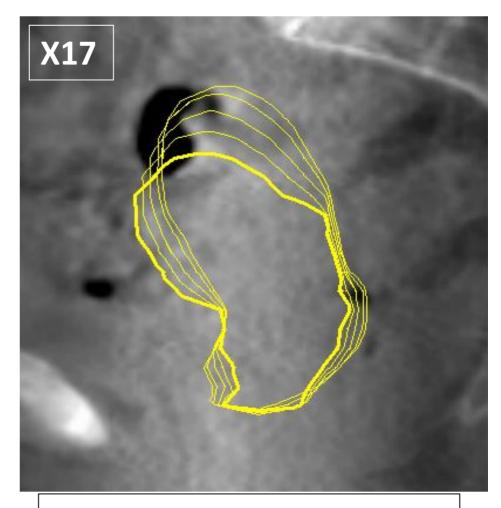




Patient 5



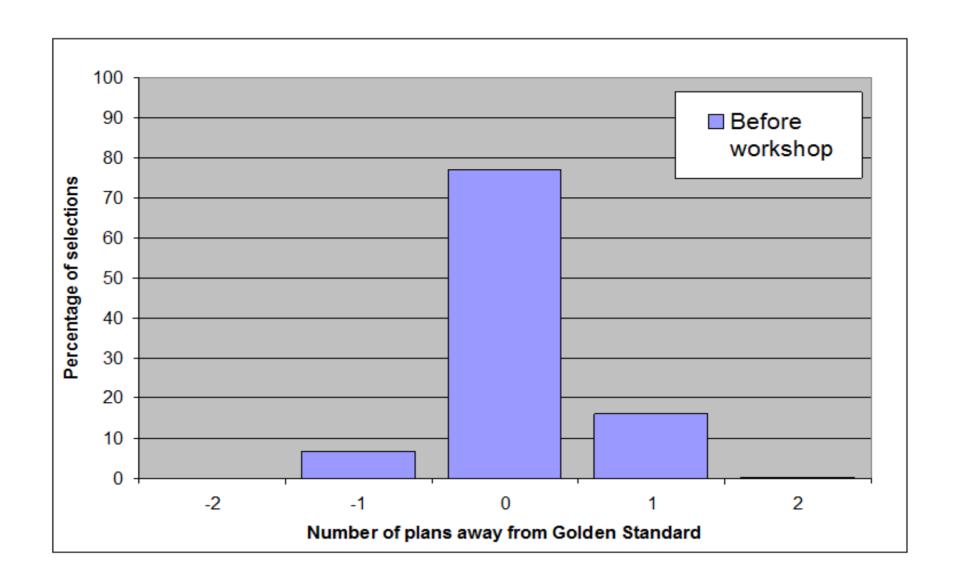
100% agreement



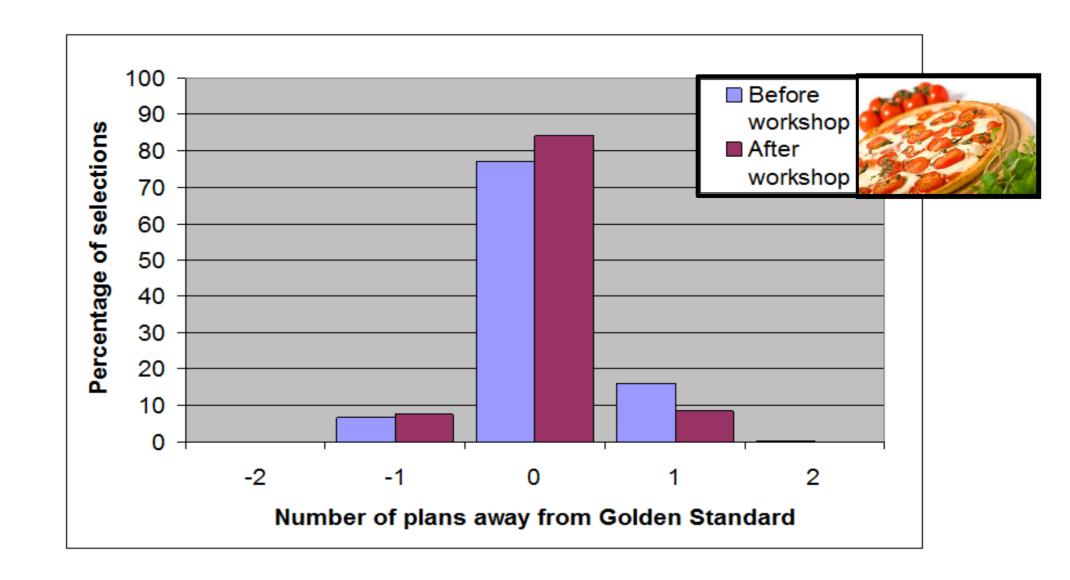
No agreement. Large anatomical change wrt planning CT



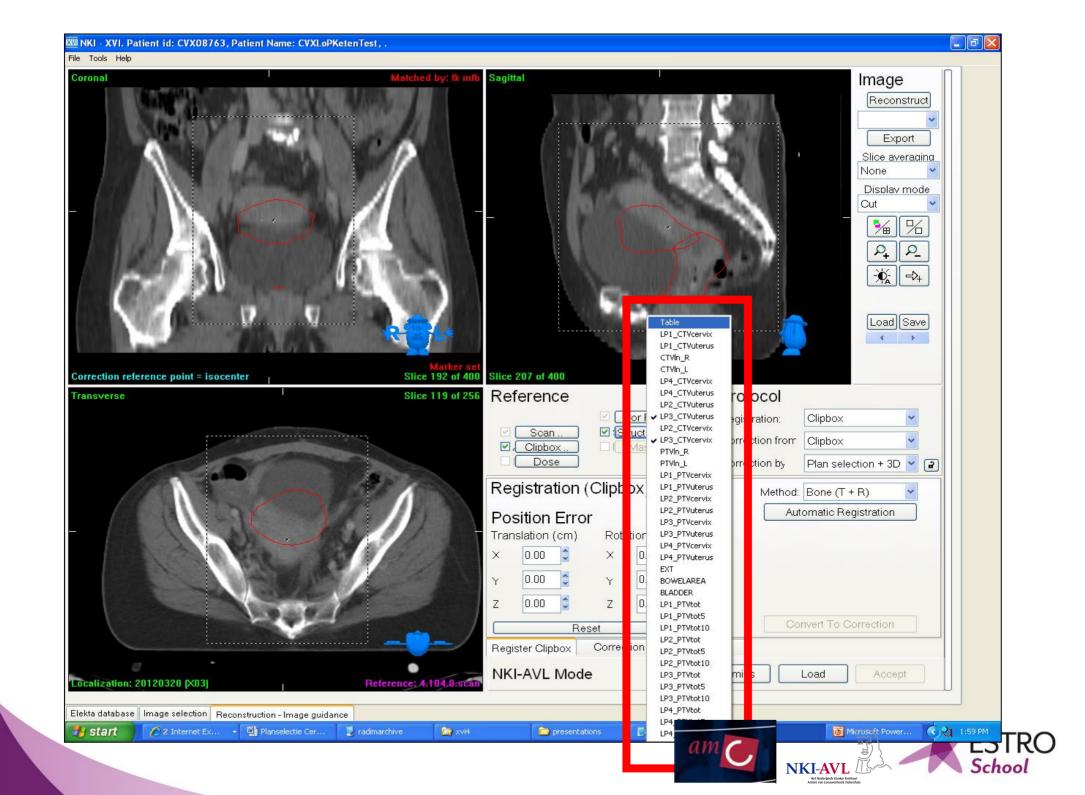


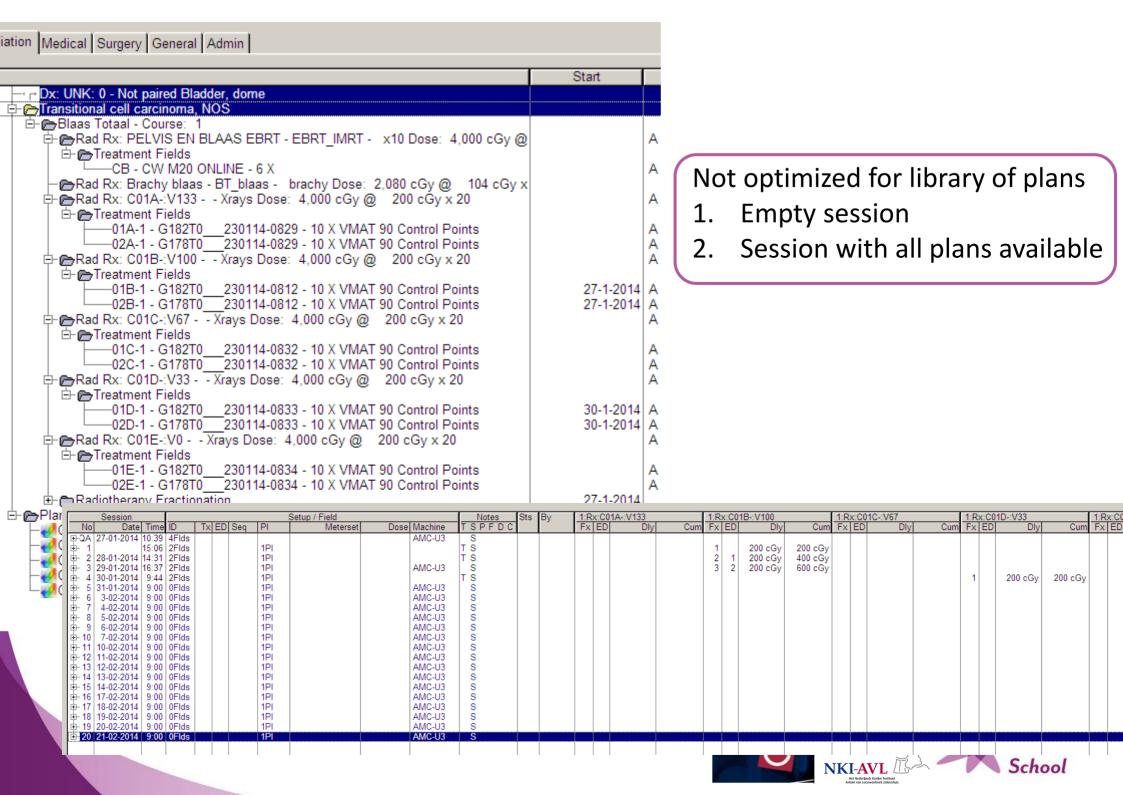


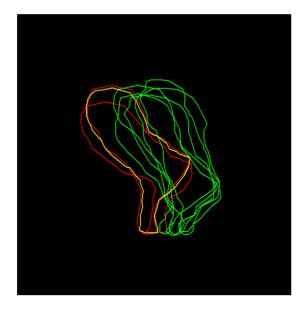


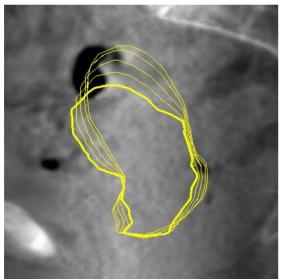












Not all anatomy change is due to bladder filling ~30% rectum filling

Anatomy change over course of treatment due to regression





ART - Cervix

	_
Patiënt naam:	
Patiënt nummer:	8890230
Course:	1
Aantal plannen	3
	3

X | 🔛 🛂 + 🖭 + | 🖚

Gegevens opslaan

Fractie	IGRT-laborant(en)	Datum	Plan	Opmerkingen
1		10 juni 2015	66-100%	
2		11 juni 2015	66-100%	
3		12 juni 2015	66-100%	
4		15 juni 2015	33-66%	krap
5		16 juni 2015	66-100%	
6		17 juni 2015	33-66%	buiten PTV
7		18 juni 2015	33-66%	op PTV
8		19 juni 2015	66-100%	krap
9		22 juni 2015	66-100%	op PTV
10	mka/jvr	23 juni 2015	33-66%	op PTV, bekkenkanteling, klieren gecovered
11	rjg/kgl/jvr	24 juni 2015	66-100%	krap binnen PTV craniaal
12	dwl mos ahk	25 juni 2015	66-100%	craniaal buiten PTV
13				
14				
15				
16				
17				
18				
19				
20				

Week	Eval.	Door	Opmerkingen
1	V	rjg	
2	Ŋ	rjg	iom LSS en EPQ volume vergroting akkoord
3			
4			

Per 13 april 2015 zijn we begonnen met planselectie voor cervixpatiënten: ART cervix. Zie voor complete werkinstructie Kwadraet 'Cervix ART'. Iedere dag moet er iemand van de IGART-groep aanwezig zijn bij planselectie.

Om inzichtelijk te maken voor het toestel welke plannen er worden gekozen graag bovenstaande tabel invullen. Als er vroeg in de behandeling 3 keer achter elkaar voor plan 0% - 33% wordt gekozen, graag in gesprek gaan met de patiënt of hierin verbetering kan worden aangebracht door meer te drinken of eerder uit te plassen. Dit graag noteren bij 'opmerkingen'.

NB: In tegenstelling tot ART-Blaas mag er bij ART-Cervix NOOIT getweakt worden!

Vragen en/of opmerkingen horen wij graag. Alvast dank.

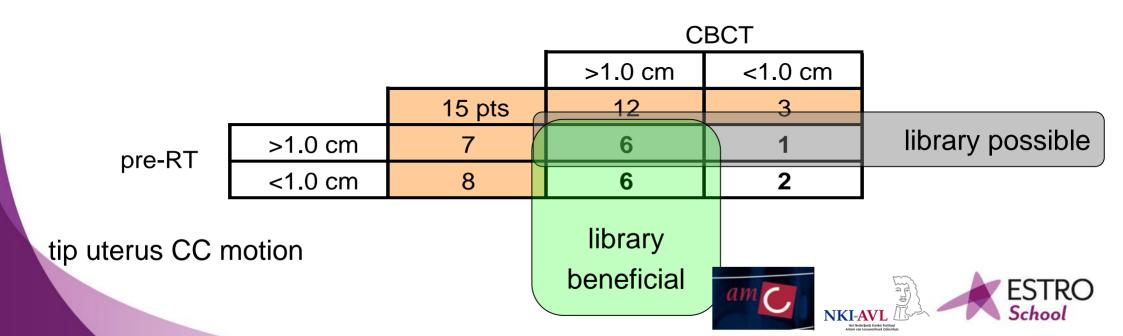
Jorrit (62594) Rianne (66857)

Gereed Bestralingsoverzicht

CT scans full and empty bladder

- 1 full bladder CT
- 1 empty bladder CT

Judgement call of the RTT at CT simulation whether bladder is "full" If not, patient needs to wait extra time and rescan.



CT scans full and empty bladder

- RTT registers CT scans and determines movement:
 - <1 cm no planselection
 - Between 1 en 2 cm: 2 plans
 - 2 cm < 4 plans
- Physician delineates target volumes on full and empty bladder CT scan
- Physicist generates intermediate structures and PTV margins
- RTT delineates OAR and generate treatment plan with VMAT on full bladder CT scan, scripting generates the intermediate plans



Planselection workflow at Linac

- 1. Registration bones
- 2. Selection of plan
- 3. Marker check
 - Trained RTTs

Demo database for training of RTTs

Workshop

Extra, not commercial @AvL/NKI

Big Brother software checks consistency of selected plan in XVI and Mosaiq plan!

Big Brother software prohibits delivery of more than 1 plan!



Evaluation / safety procedures

1 x week by IGRT-group (expert RTT)

- Selection as discussed in workshop?
- Is uterus moving as predicted on full/empty bladder CT scan (mover/non-mover)?
 - Including those patients that had no planselection due to no movement at full/empty bladder CT scans
- Are the anatomy changes still valid? Think regression



Documents...

- Patient letters for bladder instructions
- RTT instructions for full bladders
- MRI of Pet CT in full bladder? CT scan full bladder obsolete?
- Delineation instructions and interpolations
- Mosaiq
- Treatment Planning scripting
- IGRT



Discussion

Cervix planselection: a RTTs job?

(Almost) every step of the way





Summary

- Development and implementation of ART (library of plans) is
 - Departement specific
 - Protocol specific
- Development and implementation of ART is a multi-diciplinairy effort
- Because of the multi-disciplinairy character one needs to be creative, like pizza meetings
- Training is key. Invest in training as it will improve quality, but also raises awareness as to the importance of IGRT/ART!



Special thanks to

AvL/NKI
Folkert Koetsveld
Simon van Kranen
Peter Remeijer

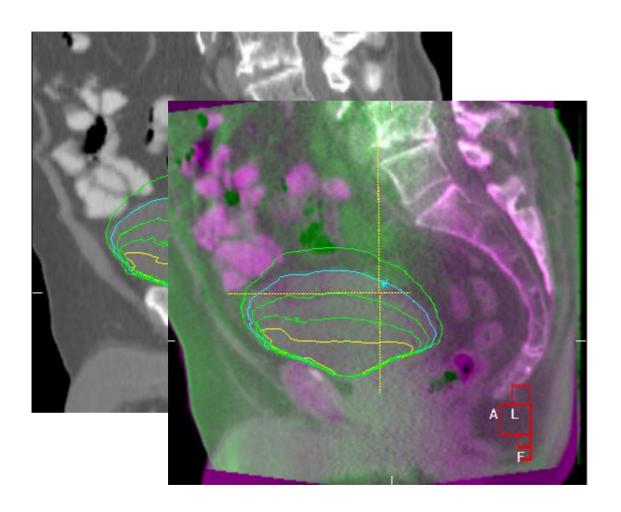








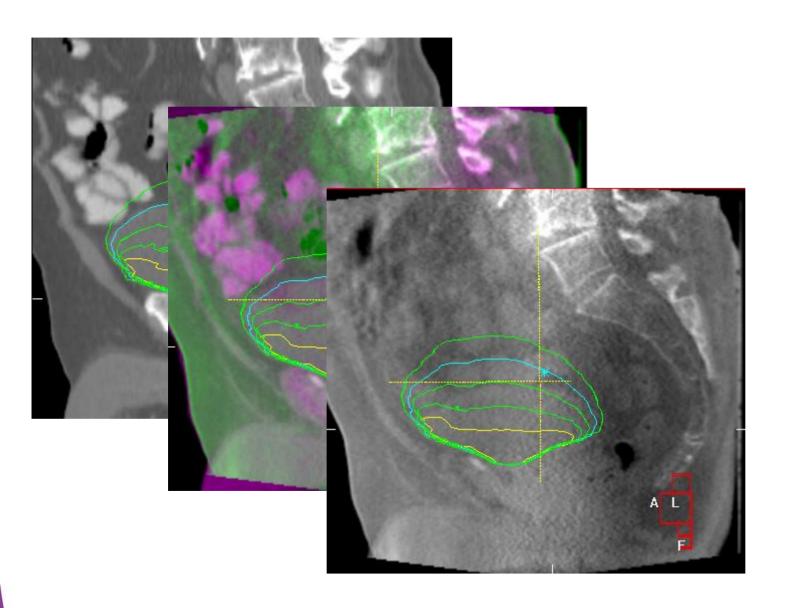








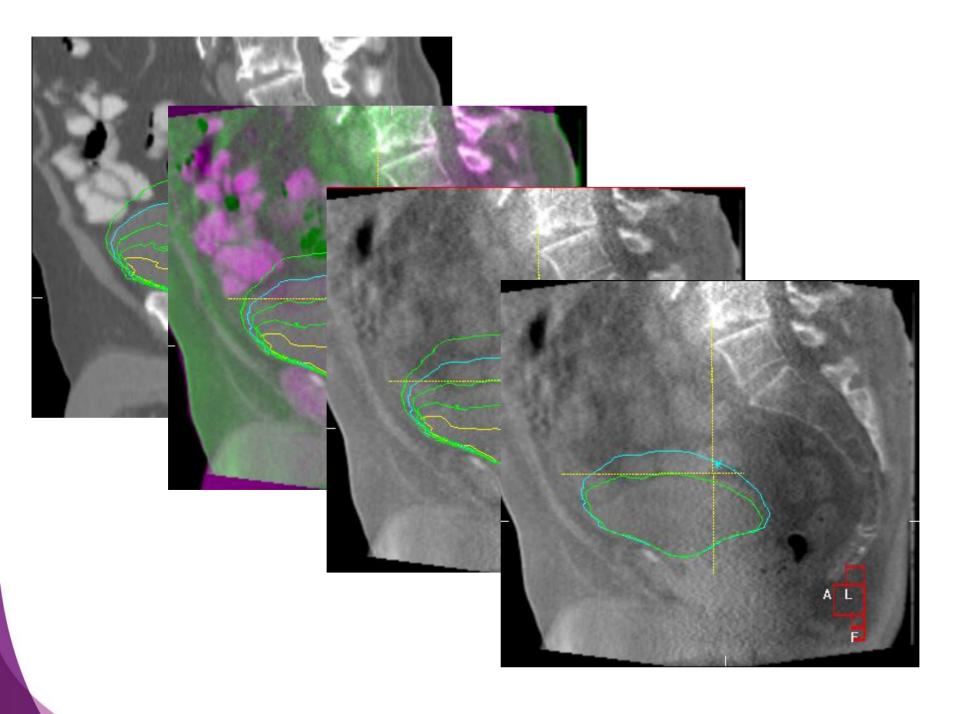








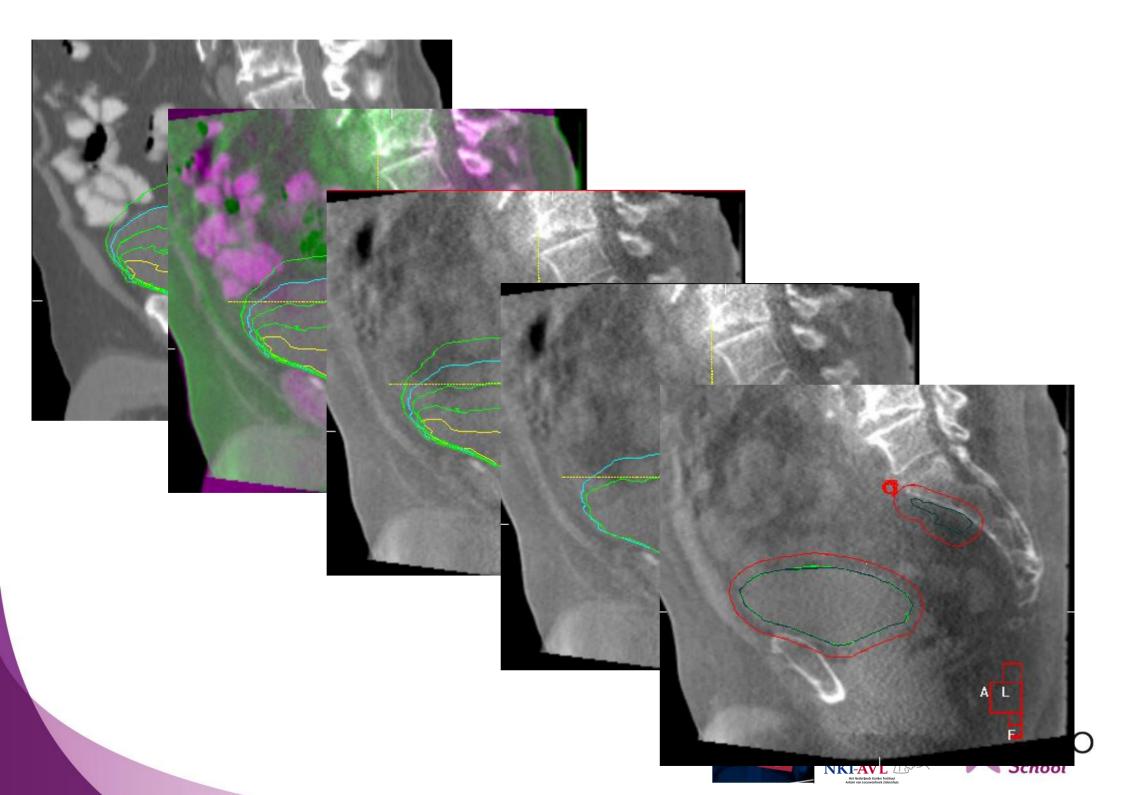


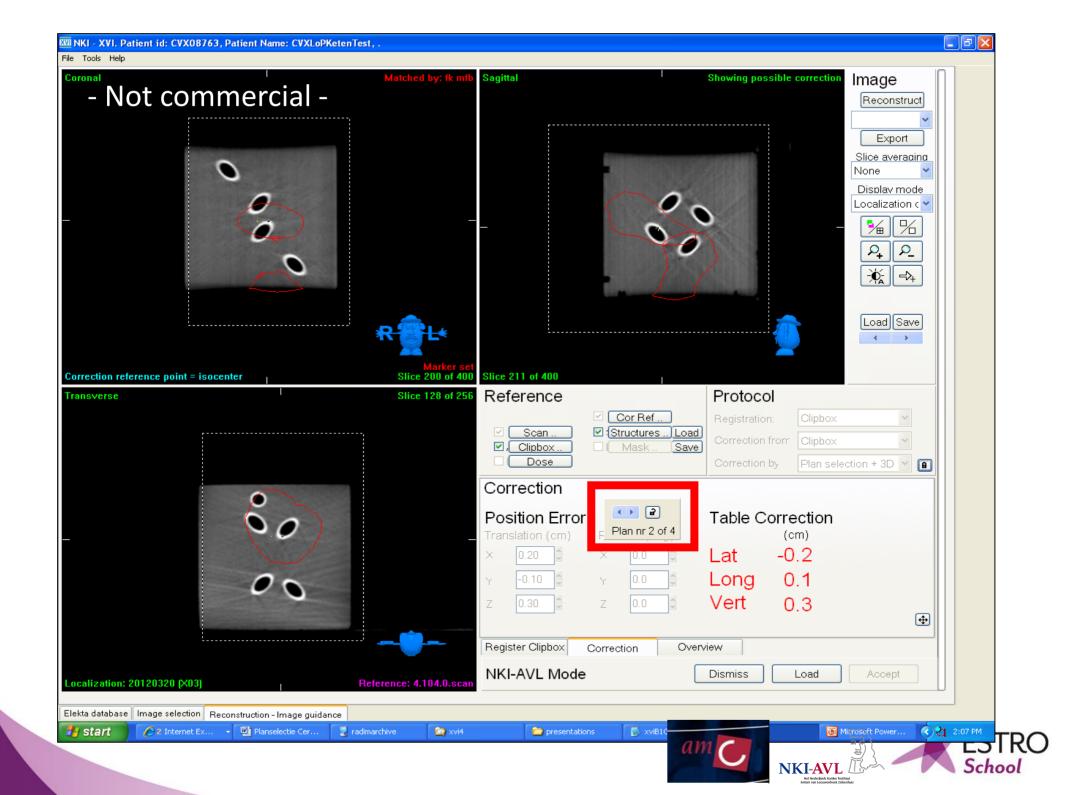


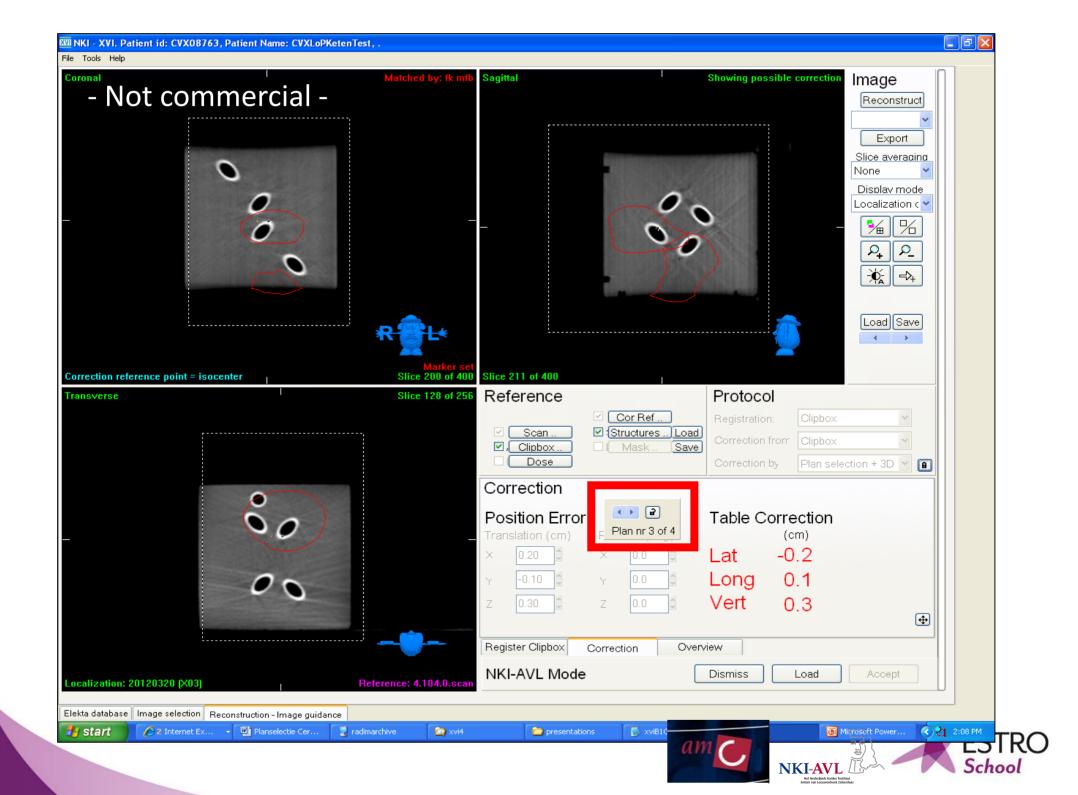


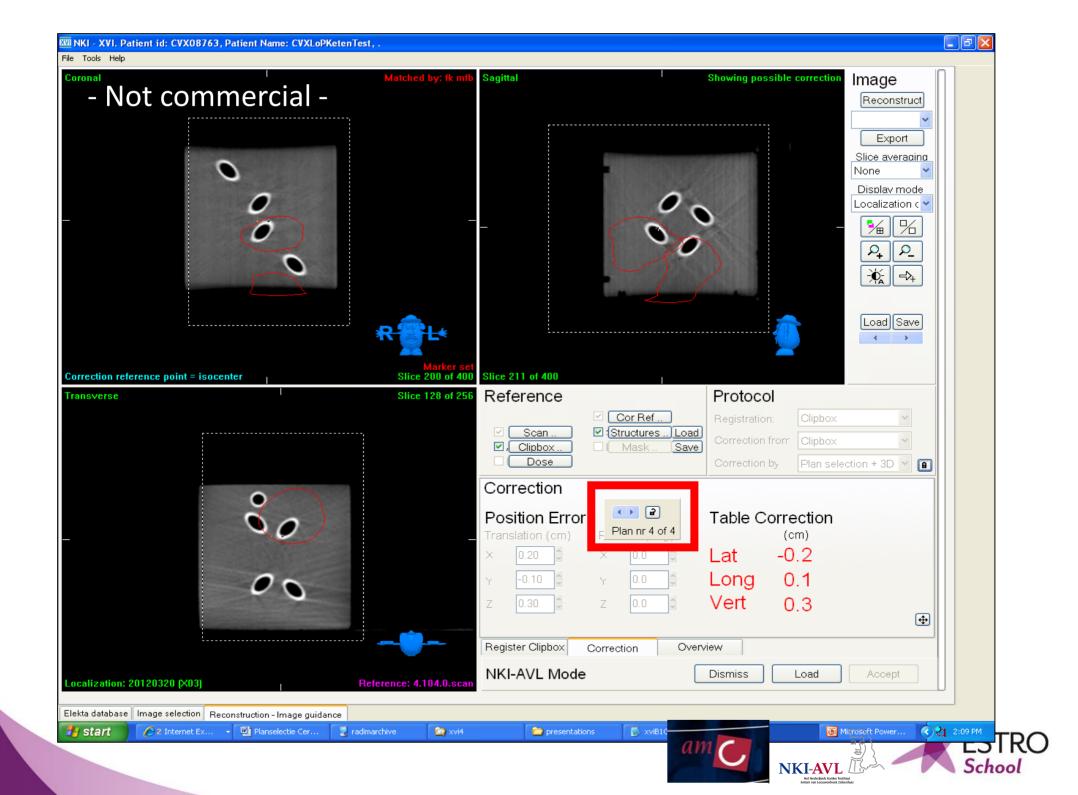


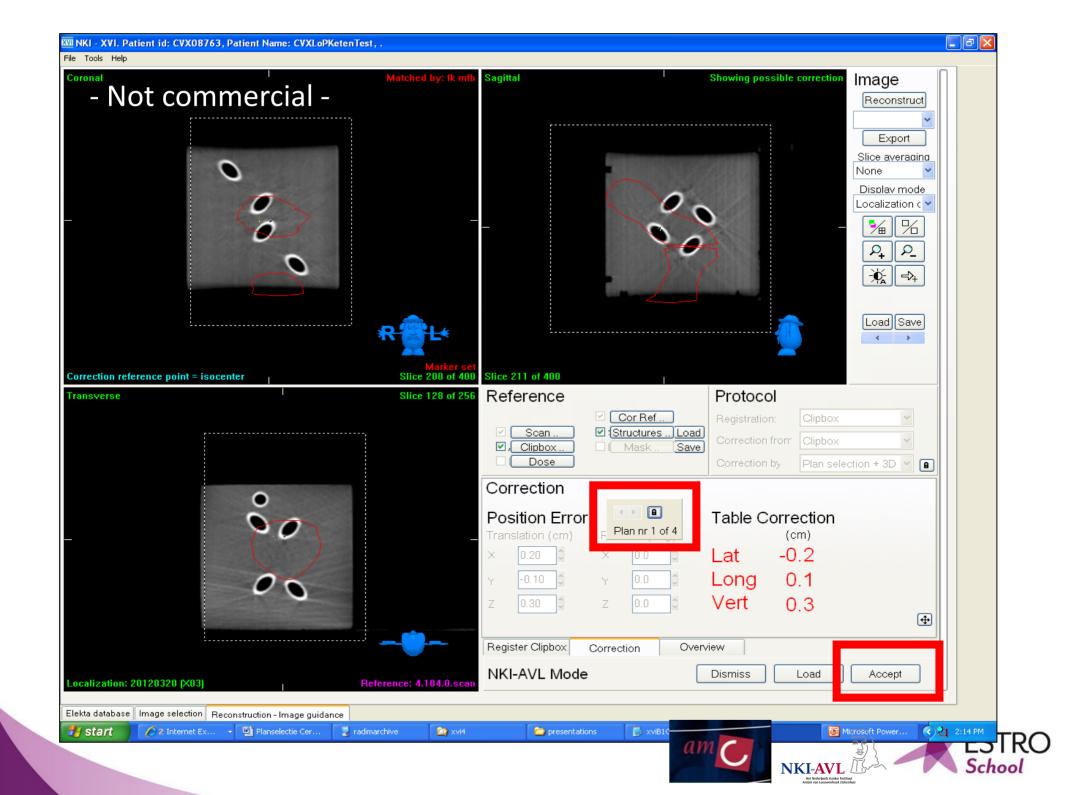


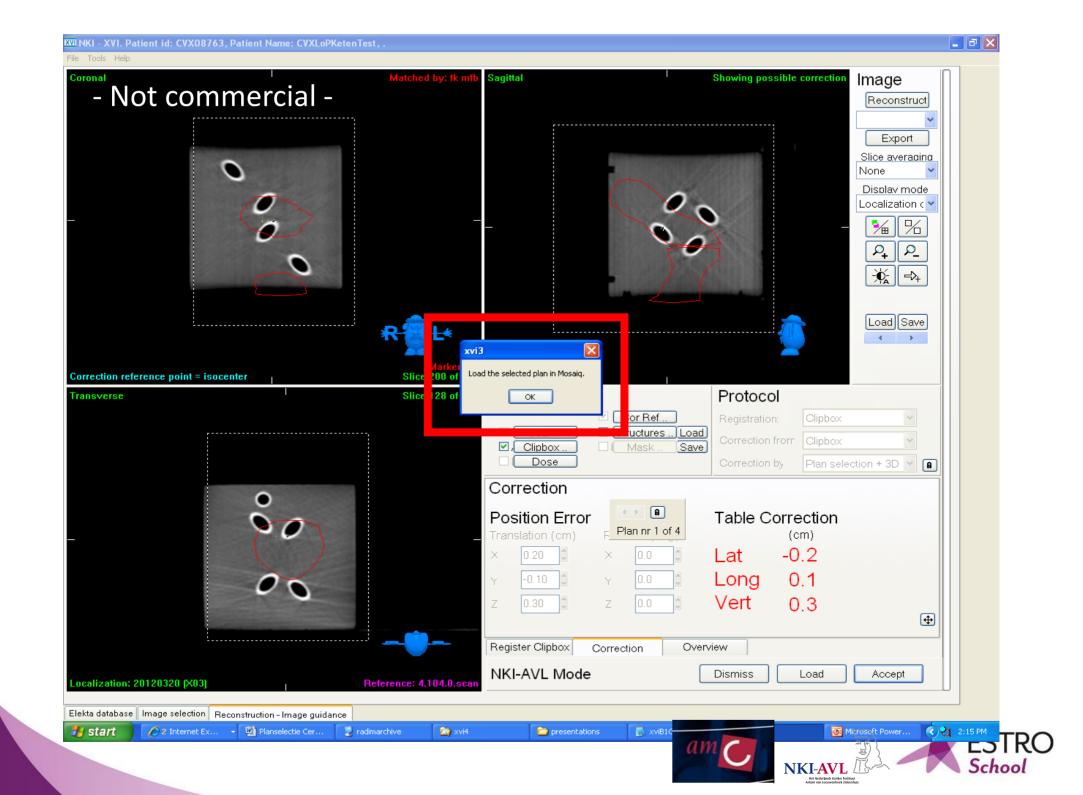














Case reports: Prostate a physicist's perspective





Mirjana Josipovic

Dept. of Oncology, Rigshospitalet & Niels Bohr Institute, University of Copenhagen Denmark

Advanced skills in modern radiotherapy June 2017



Imaging for prostate RT planning

Imaging immobilised patient in the treatment position

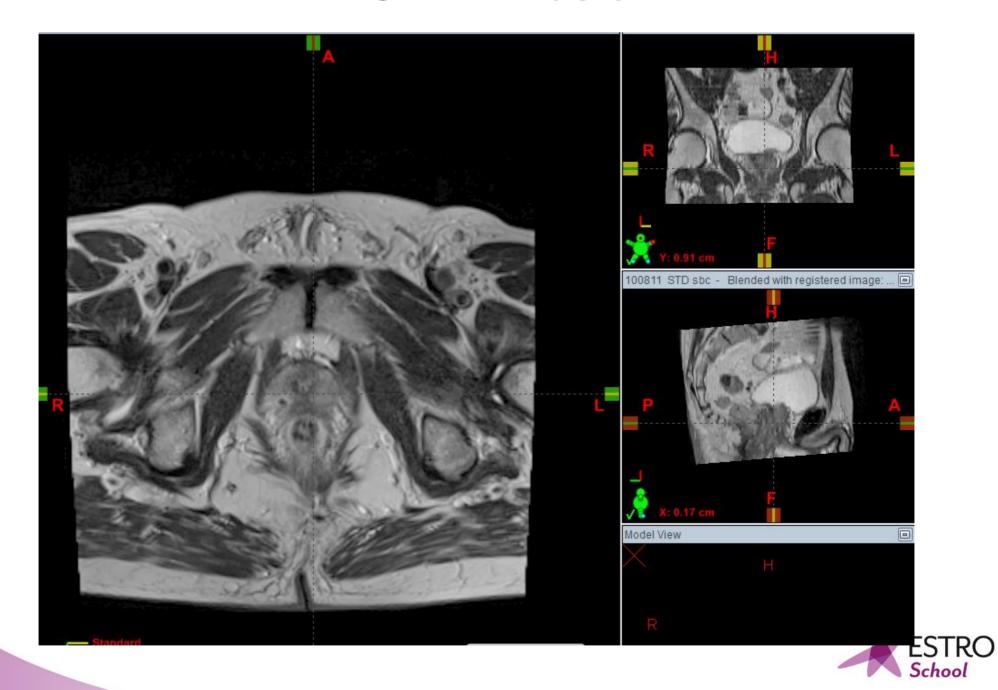
- CT scan
- MR scan
- Marker implantation into prostate gland



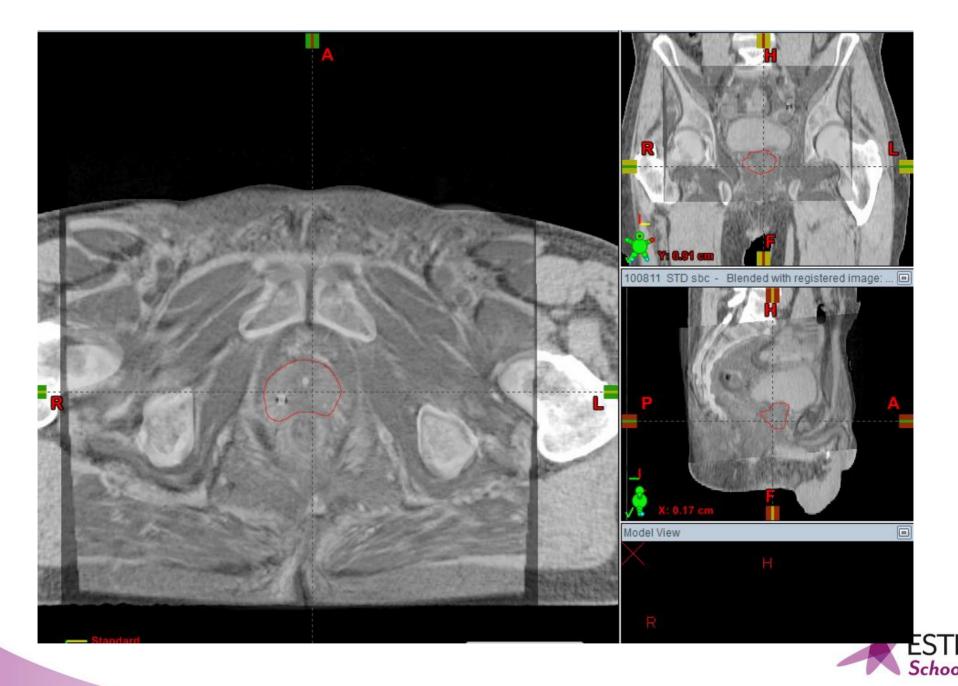
CT MR fusion



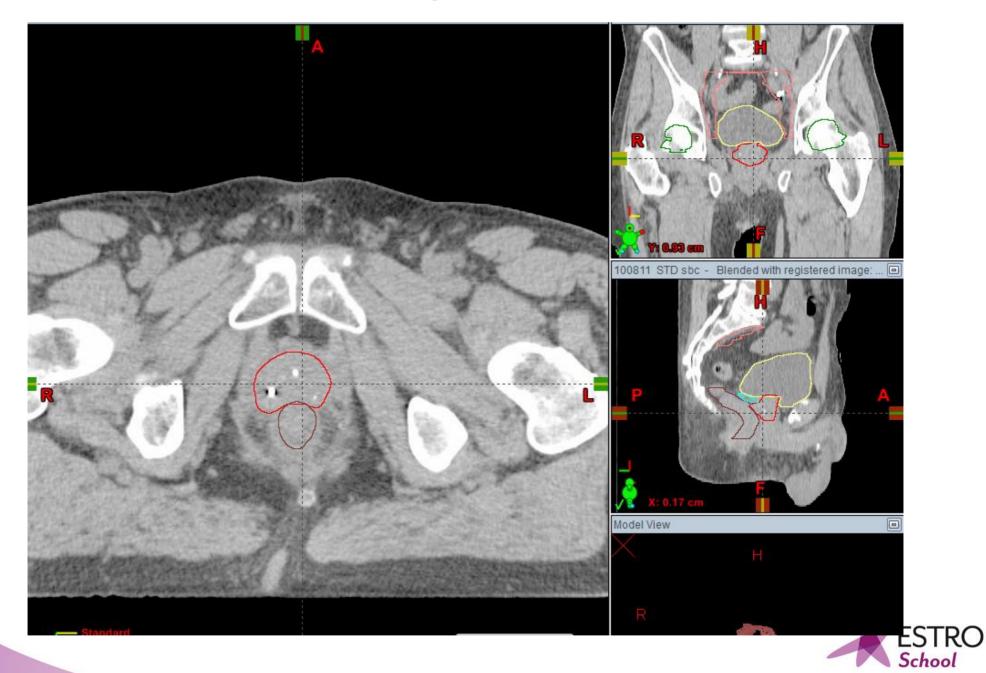
CT MR fusion



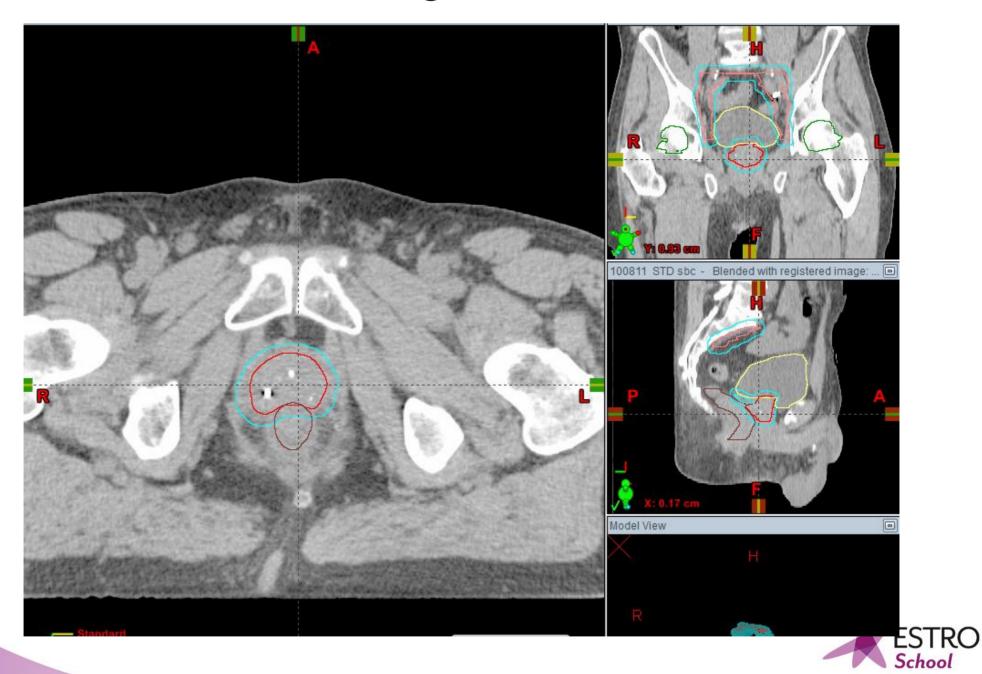
CT MR fusion

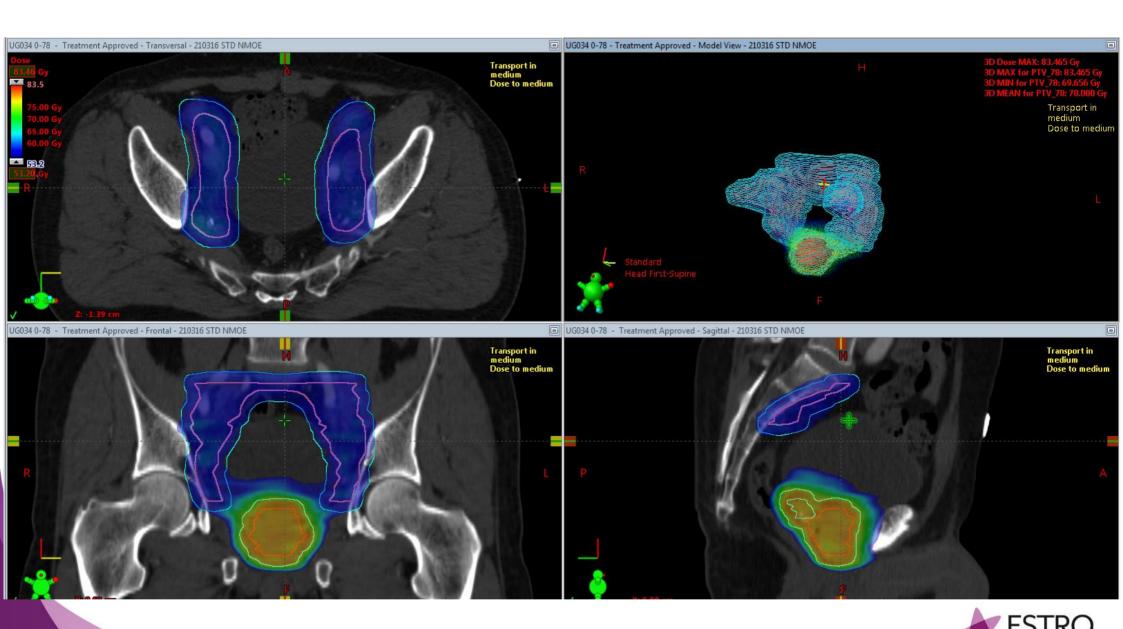


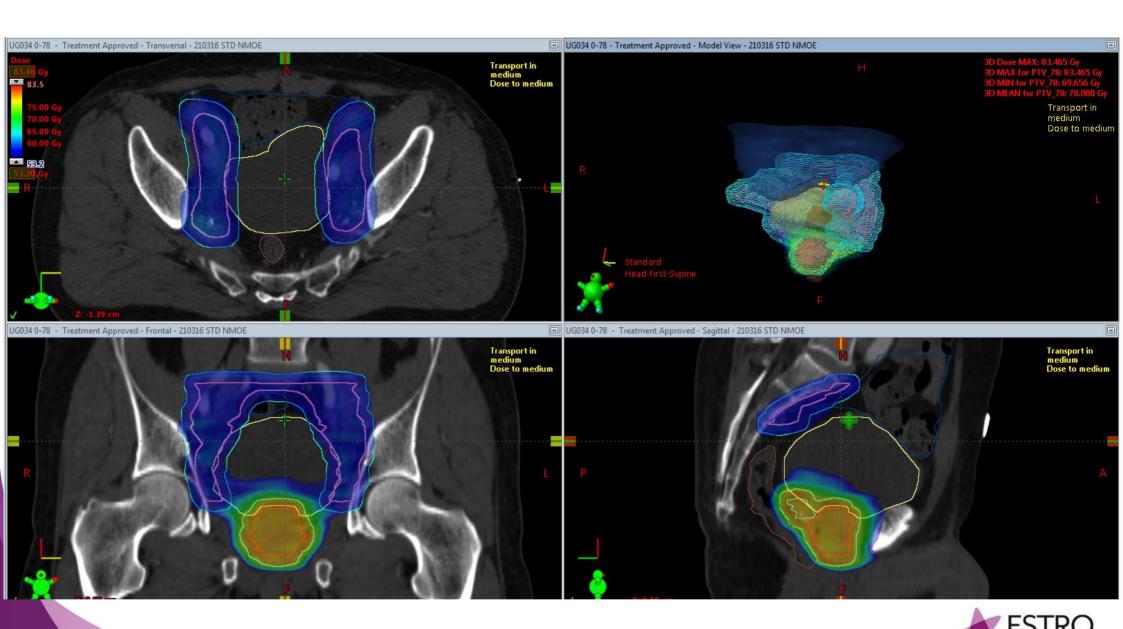
Target & OAR

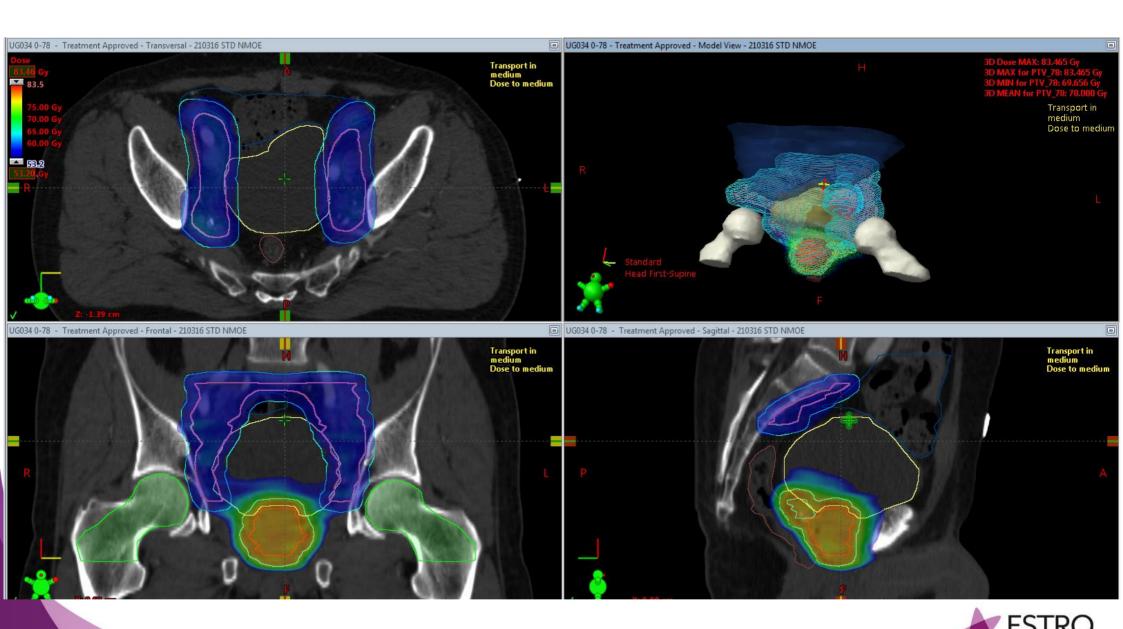


Target & OAR









Treatment planning

Copenhagen: 39 fractions of

- 2 Gy (total 78 Gy) to the prostate
- 1.49 Gy (total 58 Gy; $EQD2_{1.5} = \sim 50$ Gy) to the sem.vessicles & nodes

Seville: 28 fractions of

- 2.32 Gy (total 65 Gy; $EQD2_{1.5} = ~71$ Gy) to the prostate
- 2.14 Gy (total 60 Gy; $EQD2_{1.5} = \sim 62$ Gy) to the sem.vessicles & inv.nodes
- 1.75 Gy (total 50 Gy; $EQD2_{1.5} = \sim 37$ Gy) profilactic to the noninv.nodes

Challenge of overlap of PTV & OAR



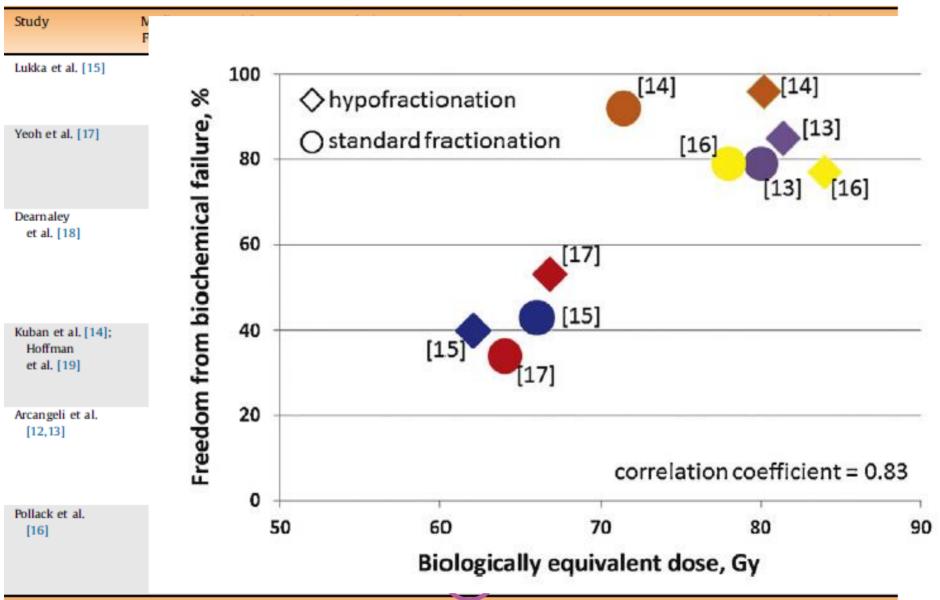
Table 1 - Phase 3 randomized trials of moderate hypofractionation for intact prostate cancer

Study	Median FU, mo	Risk, GS, or NCCN	Technique	Regimen	BED, Gy	n	Outcome	Toxicity
Lukka et al. [15]	68	60% GS ≤6 31% GS 7 9% GS 8–10	3DCRT No IGRT	5/2.5 Gy/20 fx	62	466	5 yr FFBF 40% (NS)	Gr ≥3 2% (NS)
				66 Gy/33 fx	66	470	5 yr FFBF 43%	Gr ≥3 1%
Yeoh et al. [17]	90	n.s.	2D/3DCRT No IGRT	55 Gy/20 fx	66,8	108	7.5 yr FFBF 53% (p < 0.05)	Late GU; HR: 1.58 (95% CI, 1.01-2.47) favoring hypofractionation
				64 Gy/32 fx	64	109	7.5 yr FFBF 34%	
Dearnaley et al. [18]	51	n,s,	3D/IMRT No IGRT 3–6 mo AI	57 Gy/19 fx	73 4	151	n.s.	Gr ≥2 GU 0% (NS) Gr ≥2 GI 1% (NS)
				60 Gy/20 fx	77	153		Gr ≥2 GU 2% Gr ≥2 GI 4%
				74 Gy/37 fx	74	153		Gr ≥2 GU 2% Gr ≥2 GI 4%
Kuban et al. [14]; Hoffman et al. [19]	60	28% low 71% intermediate 1% high	IMRT IGRT 21% ADT	72 Gy/30 fx	80.	102	5 yr FFBF 96% (NS)	5 yr Gr ≥2 GU 16% (NS) 5 yr Gr ≥2 GI 10% (NS)
			- 1	75.6 Gy/42 fx	71.	101	5 yr FFBF 92%	5 yr Gr ≥2 GU 17% 5 yr Gr ≥2 GI 5%
Arcangeli et al. [12,13]	70	26% GS ≤7 74% GS >7	3DCRT No IGRT 100% 9 mo ADT	62 Gy/20 fx	81 4	83	5 yr FFBF 85% (p = 0.065) *p ss for GS ≥4 + 3	3 yr Gr ≥2 GU 16% (NS) 3 yr Gr ≥2 GI 17% (NS)
				80 Gy/40 fx	80	85	5 yr FFBF 79%	3 yr Gr ≥2 GU 11% 3 yr Gr ≥2GI 14%
Pollack et al. [16]	68	34% GS ≤6 47% GS 7 19% GS 8–10	IMRT IGRT	70.2 Gy/26 fx	84	151	5 yr BCDF 23% (NS)	5 yr Gr ≥2 GU 13% (p = 0.16) 5 yr Gr ≥2 GI 9% (NS)
				.8 Gy/36 fx	78	152	5 yr BCDF 21%	5 yr Gr ≥2 GU 13% 5 yr Gr ≥2 GI 9%

3DCRT = three-dimensional conformal radiotherapy; ADT = androgen-deprivation therapy; BCDF = biochemical or clinical disease failure; BED = biologically equivalent dose, calculated to be equivalent in 2 Gy fractions using an α/β of 1.5 Gy; CI = confidence interval; FFBF = freedom from biochemical failure; FU = follow-up; fx = fractions; GI = gastrointestinal; Gr = grade; GS = Gleason score; GU = genitourinary; HR = hazard ratio; IGRT = image-guided radiation therapy; IMRT = intensity-modulated radiation therapy; NCCN = National Comprehensive Cancer Network; NS = not significant; n.s. = not stated; ss = statistically significant.

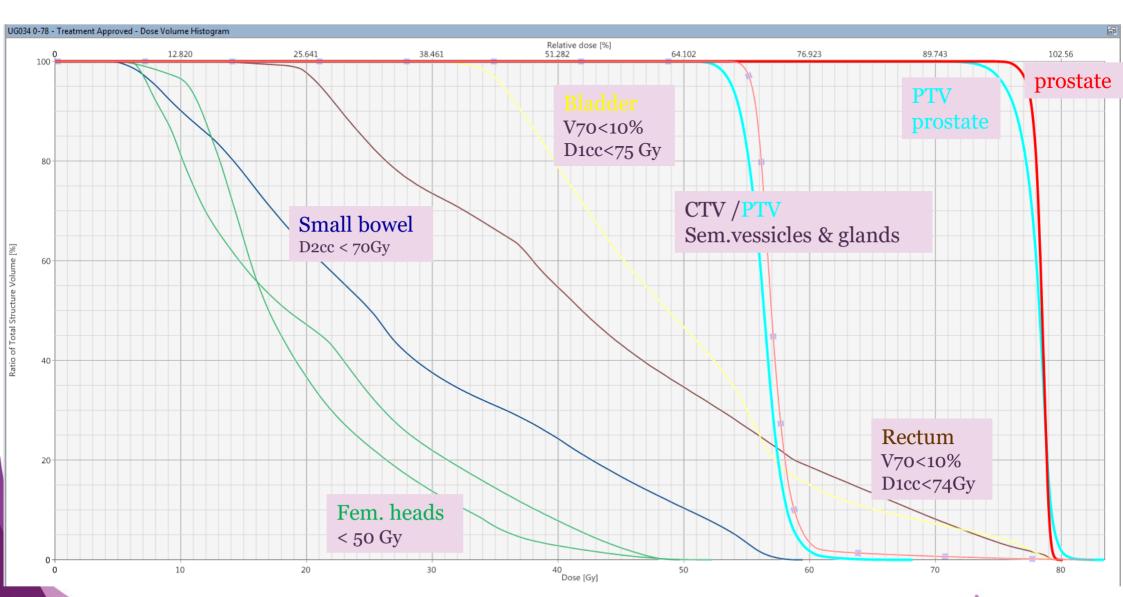


Table 1 - Phase 3 randomized trials of moderate hypofractionation for intact prostate cancer



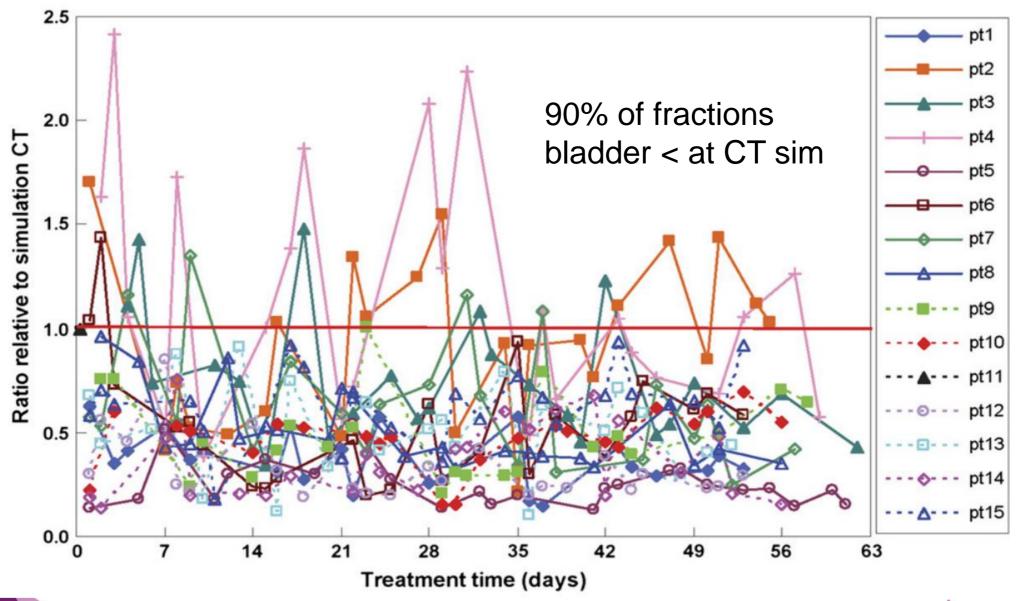
3DCRT = three-dimensional conformal radiotherapy; ADT = androgen-deprivation therapy; BCDF = biochemical or clinical disease failure; BED = biologically equivalent dose, calculated to be equivalent in 2 Gy fractions using an α/β of 1.5 Gy; CI = confidence interval; FFBF = freedom from biochemical failure; FU = follow-up; fx = fractions; GI = gastrointestinal; Gr = grade; GS = Gleason score; GU = genitourinary; HR = hazard ratio; IGRT = image-guided radiation therapy; IMRT = intensity-modulated radiation therapy; NCCN = National Comprehensive Cancer Network; NS = not significant; n.s. = not stated; ss = statistically significant.







Changes in bladder volume during the RT

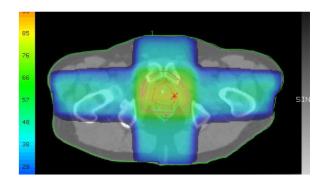




Treatment techniques

< 2005

3D conformal "box" technique



2005

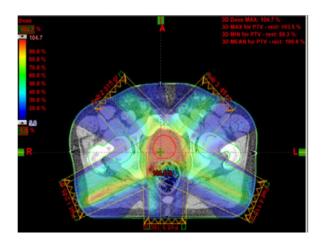
IMRT

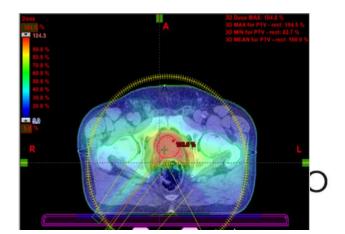
- Gold seeds & daily IGRT
- Margins ↓

2008

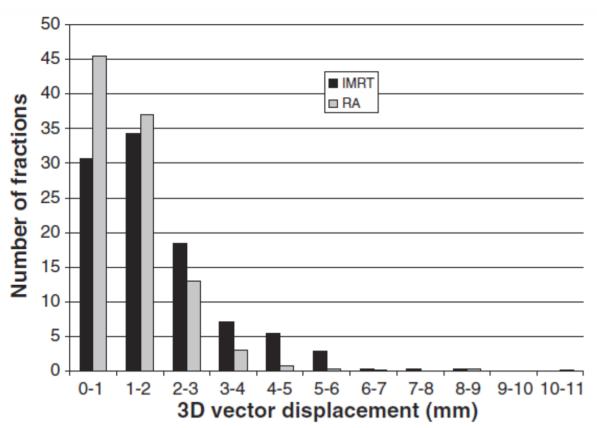
VMAT

- Faster treatment (1 arc = 1 min)
- Margins ↓





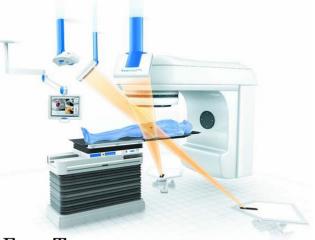
Treatment delivery time



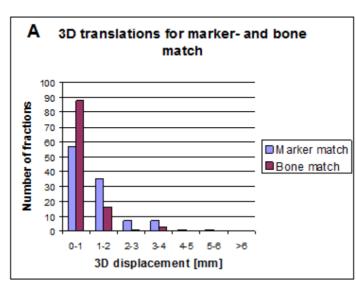
	IMRT	VMAT
Treatment delivery time	4.9 min	1.1 min
Intra-fractional prostate displacement > 3mm	16.7%	4.7%



Prostate rotation

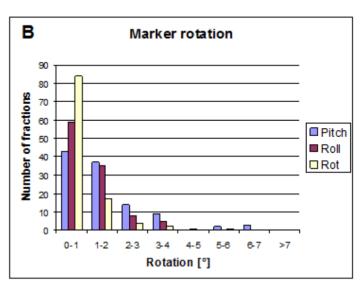


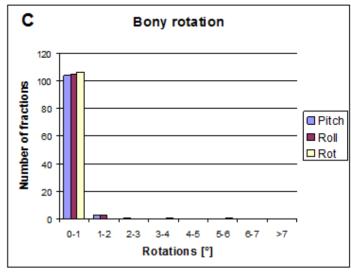
Exac Trac with robotic couch



After the 6D correction, the deviation in prostate is larger than deviation in patient position, for both translation and rotation

Courtesy of JS Rydhög







IGRT for prostate cancer: RTT perspective

Martijn Kamphuis MSc MBA Research Radiation Therapist IGRT

Department of Radiotherapy @ AMC Amsterdam, the Netherlands



Content

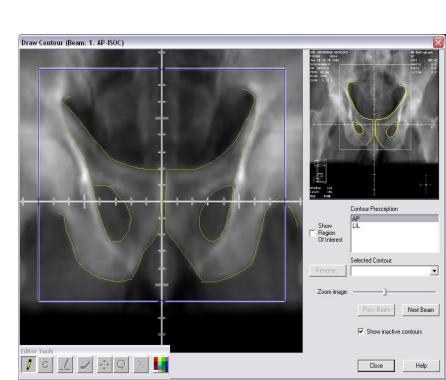
- Prostate IGRT in general
 - Offline bony anatomy matching
 - Offline marker registration using fiducial markers and PI
 - Online marker registration using fiducial markers Portal or static kV imaging
 - Online marker registration using Conebeam-CT

• IGRT for this challenging case ©



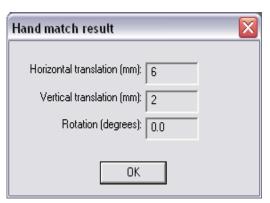
Offline/Online bony anatomy matching

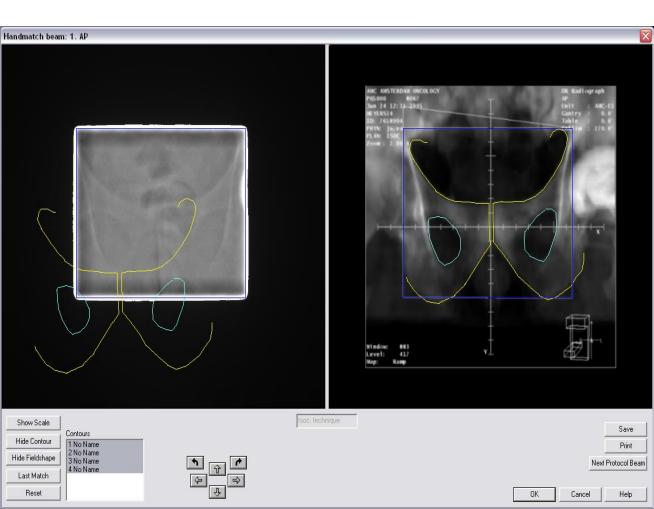
- Create an image with sufficient data
- Draw contours (templates) in reference images
- Contours should have a proper correlation with target
 - E.g. no trochantor or femur
- Produce guidelines!



Offline/Online bony anatomy matching

- Field edge match
- Match PIs





Offline marker registration using fiducial markers



Problem/challenge

 Displacement of bony anatomy does not (always) represent displacement of target



Van Herk et al.



Fiducial markers

- Displacement of bony anatomy does not (always) represent displacement of target
 - Neederveen et al. 2003: prostate cancer

	LR (mm)			AP (mm)			CC (mm)		
	marker	bone	mk. rel. bone	marker	bone	mk. rel. bone	marker	bone	mk. rel. bone
Mean	0.0	0.0	0.0	-1.0	-1.0	0.0	1.1	0.1	10
Σ	2.4	2.1	1.0	4.4	4.4	2.3	3.7	2.1	4.1
σ	2.1	1.8	0.8	3.4	2.2	2.4	2.7	1.7	2.4

Standard deviations for the systematic and random case that do differ significantly are printed in bold.

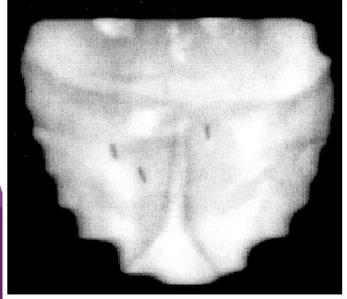
➤ For 6 out of 23 patients → increase of systematic error after correction based on bony anatomy!!

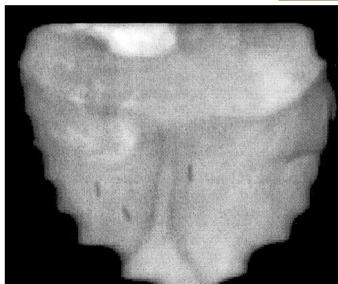


Fiducial markers: offline

Based on Van der Heide et al. 2007:

- > 5 field IMRT treatment
- > Daily offline imaging:
 - Treatment field: 40, 180 and 320 degrees
 - SAL (α =8, N=4)
 - Threshold SAL= α/\sqrt{N}
- Limited (radiation) fields adequate
 - No additional dose!







Fiducial markers: offline

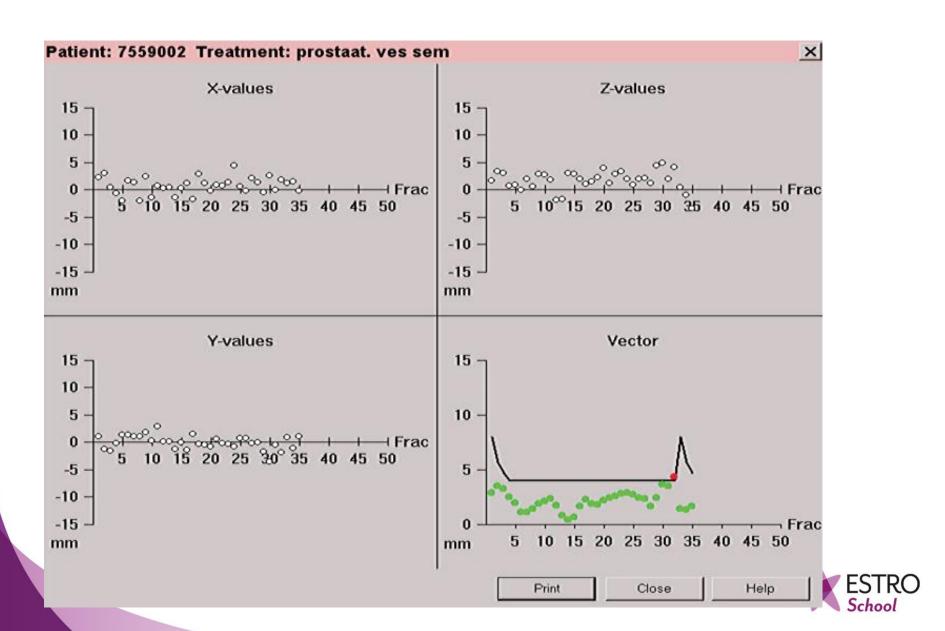
- Successfull reduction of systematic error!
 - Without applying a correction protocol, the systematic errors (Σ) are:
 - 4.8, 2.2 and 2.9 mm in the vertical, lateral and longitudinal directions
 - > The SAL protocol
 - 0.7, 0.8 and 0.8 mm, respectively.
 - Random position variations are not reduced in an off-line correction protocol



Online fiducial marker registration



Food for thought!



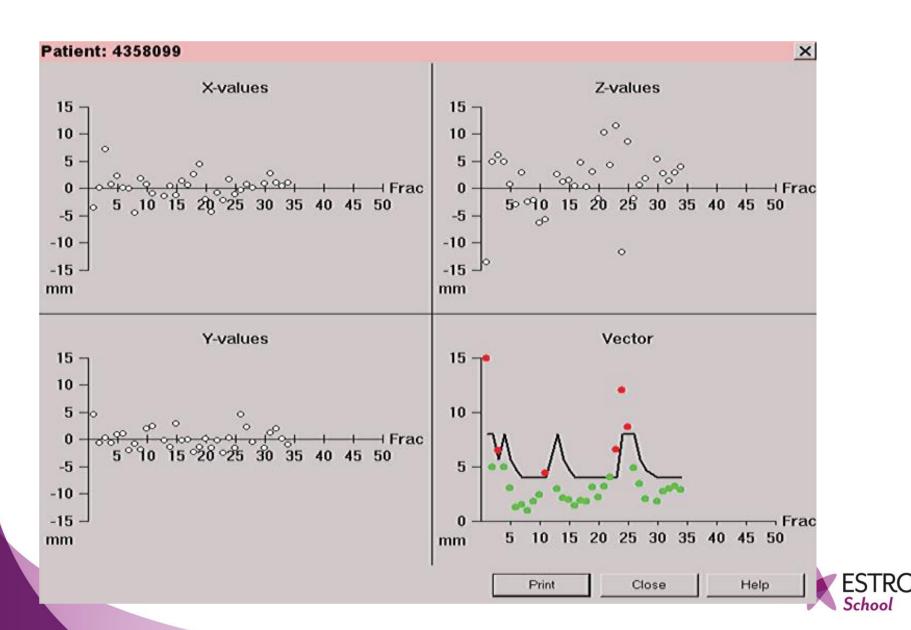
Would you like to treat a patient like this offline?

A. Yes

B. No



Food for thought!



Would you like to treat a patient like this offline?

A. Yes

B. No



Online Position Verification

- To reduce random error:
 - Online position verification is needed
- Different methods available
 - > Two dedicated EPI field, e.g. 40 and 320 degrees
 - Correction for imaging dose necessary
 - Stereo Graphic Targeting
 - MV and kV together
 - Correction for imaging dose necessary
 - > Two kV images
 - With CBCT or OBI
 - With ExacTrack system



Offline vs Online

		Results (mm)		
		X	Υ	Z
Offline	Sys. error	0.8	0.8	0.7
	Random error	2.3	2.5	4.0
Online	Sys. error	0.8	0.6	0.9
	Random error	1.0	1.0	1.2



Online Position Verification

Online procedure

Random error minimalized

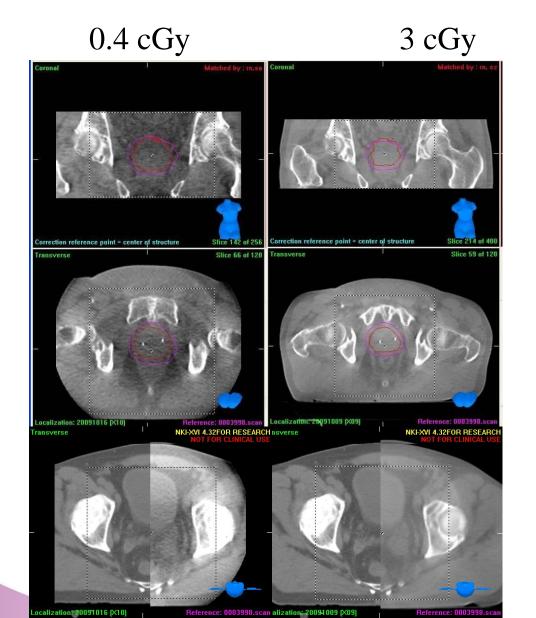
Enables **limited** margin reduction!

3.3	1.1	3.6
0.0	0.0	0.0
-2.7	-6.6	-2.7
6.6	7.2	7.1
8.0	7.8	8.7
6.8	6,6	7.5
	0.0 -2.7 6.6	0.0 0.0 -2.7 -6.6 6.6 7.2 8.0 7.8

breathing	b	0.0	0.0	0.0
scalar	a – β + σ_p	-2.7	-6.6	-2.7
CTY-PTY marge (mm)		6.2	6.2	6.1
Eenvoudige formule van Herk: 2.5°SIGMA	+0.7°sigma	7.1	6.8	7.0
Formule Stroom: 2.0°SIGMA	+0.7°sigma	5.9	5.6	5.8

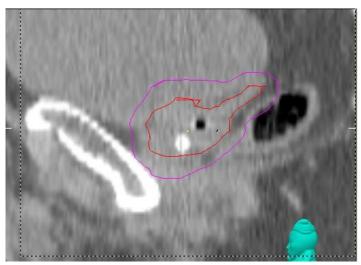


Online marker registration using CBCT



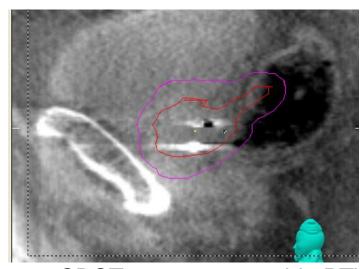


ConeBeam CT: soft tissue information



Red = Prostate + sem.ves.

Purple = PTV



CBCT : sem.ves outside PTV

Acknowledgements NKI/AvL



Many ways to Rome!

Method	Margin (AMC)	Extra <u>treatment</u> time	Imaging dose	Corretable?	Relevant anatomical information
Bone match	10 mm	2-3 minutes	(3cGY*2) High	Possible	-
Offline fiducial PI	8 mm	0 minutes	No	-	+
Online fiducial PI	(7 mm	1-3 minutes	Very low (kV) to high (MV)	Correctable in case of PI	+
Online CBCT	7 mm	1-3 minutes	0.4-3.5 cGy/scan	Partly	+++

If there is a balance with the used margin:

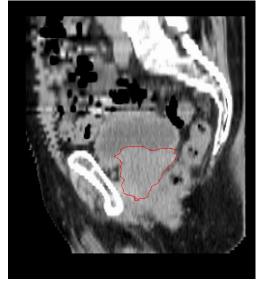
- 1. LC is about the same for the all different procedures
- 2. Toxicity probably lowest with online IGRT



The N1 case: SIB approach

Challenge: Independent moving targets

- > Lymph nodes
 - Correlate nicely with bony anatomy
- > Prostate
 - Doesn't correlate with bony anatomy



Van Herk et al.



Option 1: solved by margins (AMC)

- Use guidelines for delineation: e.g. Taylor nodes!
 Depending on correction protocol symple optimal margins
 E.g. AMC offline eNAL positive protocol 8 mm, 7 mm and 10mm for X movime arection respectively
 If ,for the night movime arection respectively
 If ,for the night movime are used
 If aware of old use 1 cm isotropic margin

 - Not optimal concerning toxicity



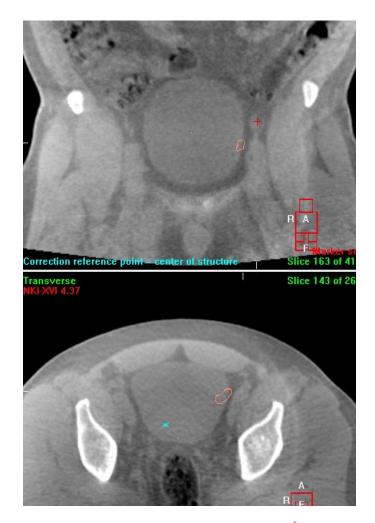
In practice large displacements were found in different cases

Correction for displacement of individual lymph nodes not always possible:

multiple targets involved

Margins can be a solution

 But first quantify the problem



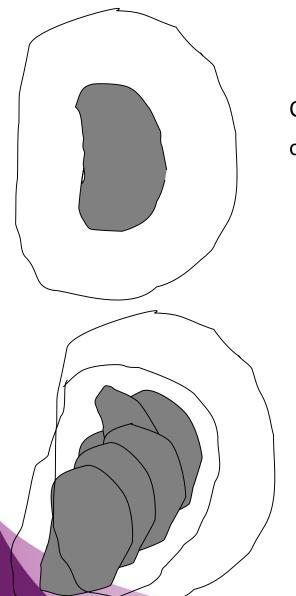


Option 2: Adaptive procedure

- Major problem:
 - "uncorrectable" systematic error between lymph nodes and prostate match
 - Peter's rules: deformations can not be corrected with table corrections



Bladder: Focal adaptive margin strategy



Conventional focal boost technique: one initial tumor position plus 2 cm margin

Adaptive margin strategy:

5 CT scans during first week of RT

Delineate 6 tumor positions plus 1 cm margin

- 40% boostvolume reduction (pos et al 06)
- less geografical missers



Thank you for your attention!



Case report: Breast

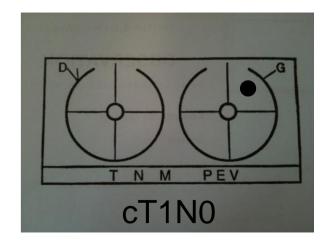


Sofia Rivera, M.D.
Radiation Oncology Department
Gustave Roussy
Villejuif, France



Advanced skills in modern radiotherapy
June 2017

Clinical case

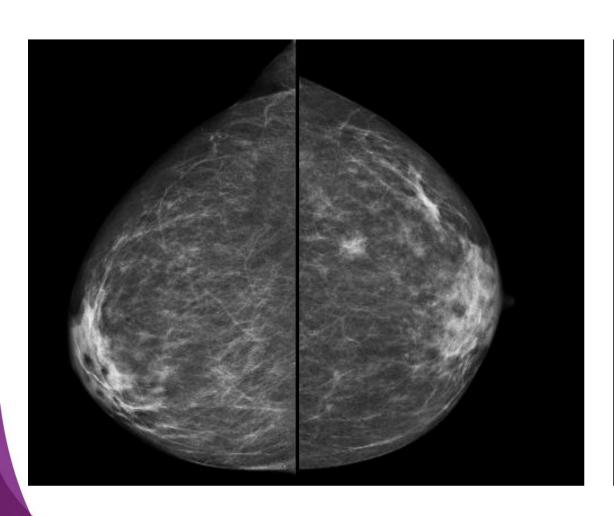


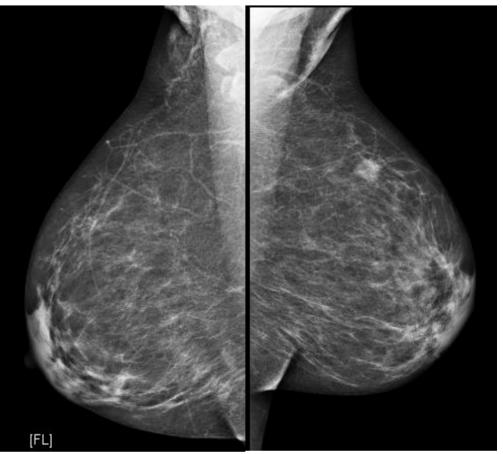
- •72 years old female patient referred by her GP after palpation of a supra areolar hard mass of the left breast external upper quadrant measuring 1cm with no axillary or supraclavicular palpable node (breast cup: 95 D)
- •Retired, yoga teacher, autonomous, living in an individual house with 5 cats
- •Medical history of hypertension, diabetes and ischemic cardiopathy



Mammograms + US





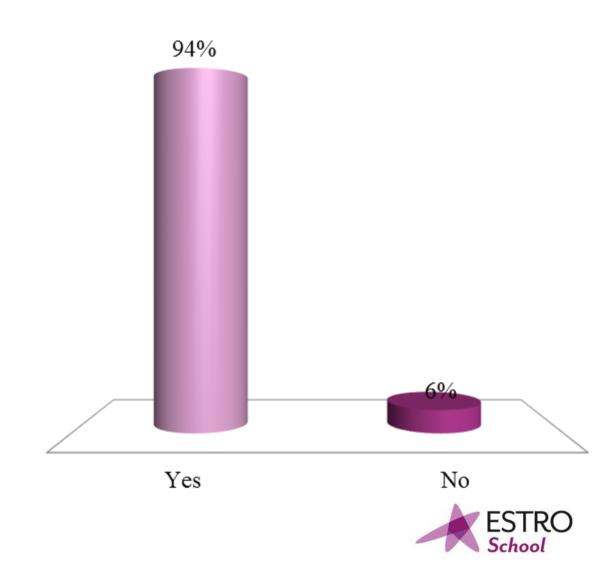




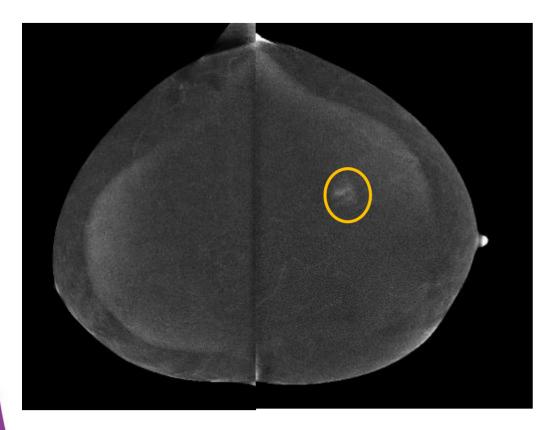
Do you see where the lesion is?

A. Yes

B. No



Angio mammography







Clinical case

- •Imaging: confirmation of a single lesion without any suspicious lymph node
- •Biopsy: Infiltrating ductal carcinoma, ER: 90%, PR: 80%, HER2-Ki67: 2%, grade I
- •Lumpectomy + sentinel lymph node procedure: pT1cNo in complete resection
- •Adjuvant radiotherapy followed by hormonotherapy for 5 years



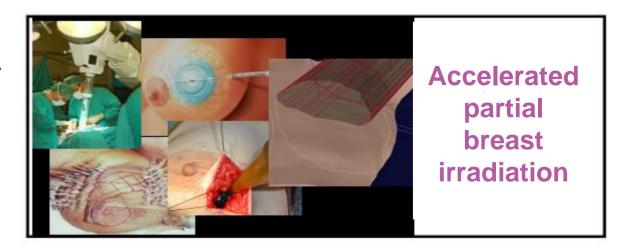
Therapeutic strategy: Which radiotherapy?

Two changing practice concepts have modified the standard whole breast irradiation 50Gy +/- boost





Whelan NEJM 2010; START A and B Lancet Oncol 2008



Vaidva Lancet 2010; Bourgier IJROBP 2010; Lemanski IJROBP 2010; Taghian IJROBP 2005; Polgar IJROBP 2004; Vicini IJROBP 2003; Formenti IJROBP 2003;

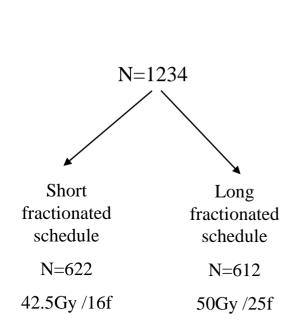


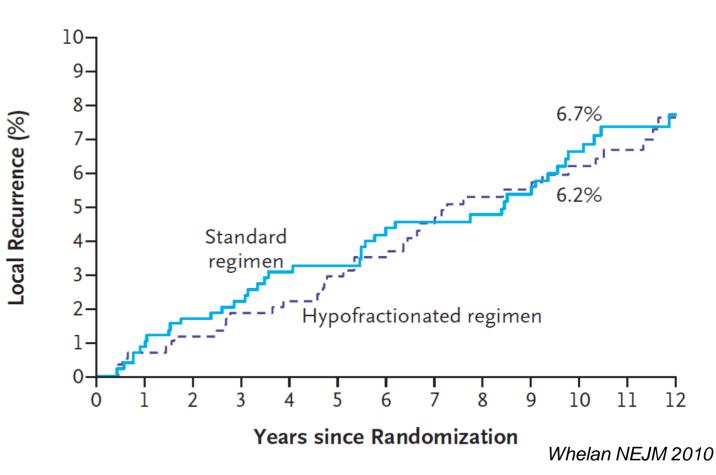
ORIGINAL ARTICLE

Whole breast irradiation

Long-Term Results of <u>Hypofractionated</u> Radiation Therapy for Breast Cancer

Timothy J. Whelan, B.M., B.Ch., Jean-Philippe Pignol, M.D., Mark N. Levine, M.D.,







Whole breast irradiation

START A
2236 patients

50 Gy/25 fractions/ 5 weeks

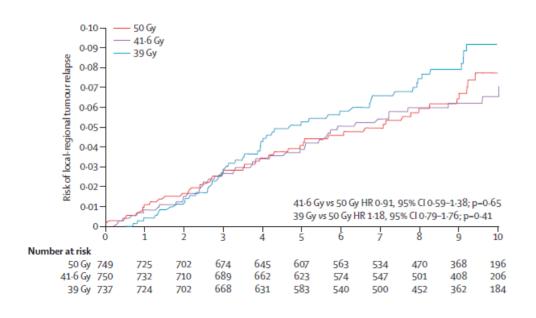
41.6 Gy/13 fractions/ 5 weeks

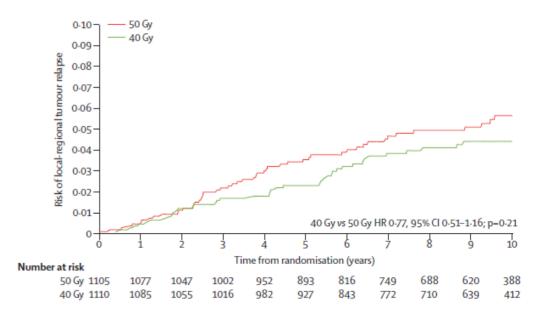
39 Gy/13 fractions/ 5 weeks



50 Gy/25 fractions/ 5 weeks

40 Gy/15 fractions/3 weeks





Median follow up = 9,3 yrs LRR-10y (50Gy) : 7,4% [5,5-10] Median follow up = 9,9 yrs LRR-10y (50Gy) : 5,5% [4.2-7,2]

JS Haviland; Lancet Oncol 2013



Partial breast irradiation indication guidelines

ESTRO

- >50 years
- IDC, mucinous, tubular, medullary, and colloid cc.
- Associated LCIS allowed but not DCIS
- Any grade, ER, PR
- pT1–2 (\leq 30 mm)
- Negative surgical margins (≥2 mm)
- Unicentric, Unifocal
- pN0 (by SLNB or ALND)

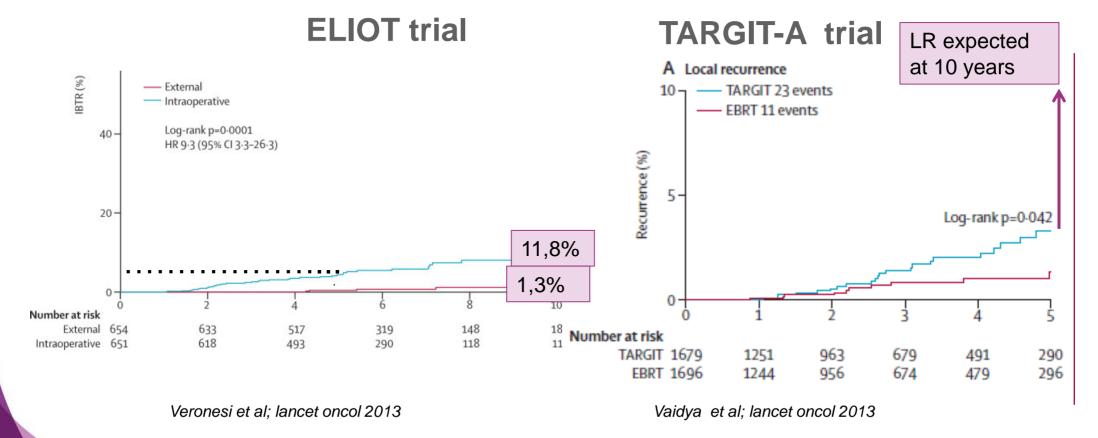
ASTRO

- \geq 60 years
- Invasive ductal or other favorable subtypes
- Pure DCIS not allowed
- ER status positive
- $pT1 : \leq 2 \text{ cm}$
- Negative surgical margins by at least 2 mm
- Unicentric only, Clinically unifocal with total size ≤2.0 cm
- pN0 (i^-, i^+) (by SLNB or ALND)



Intraoperative Partial breast versus whole breast irradiation

•Ipsilateral breast recurrence







Contents lists available at SciVerse ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



Special commentary

Has partial breast irradiation by IORT or brachytherapy been prematurely introduced into the clinic?

Harry Bartelink a,*, Celine Bourgier b, Paula Elkhuizen a

^a Netherlands Cancer Institute, The Netherlands; ^b Institut Gustave Roussy, France

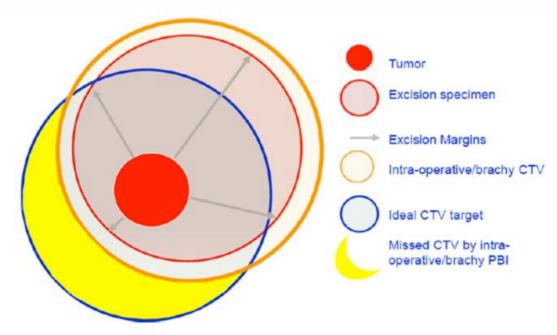


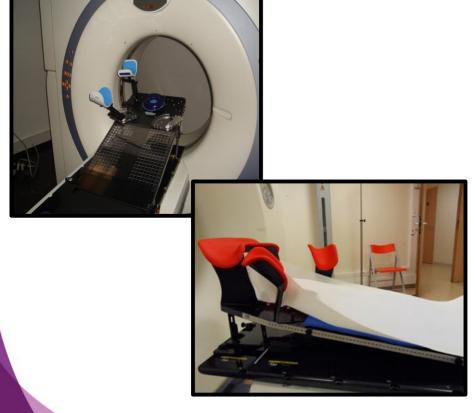
Fig. 1. Breast tumors are often eccentric located with highest risk of residual tumor in the region of the narrowest resection margin, therefore CTV by brachy or IORT is not covered.

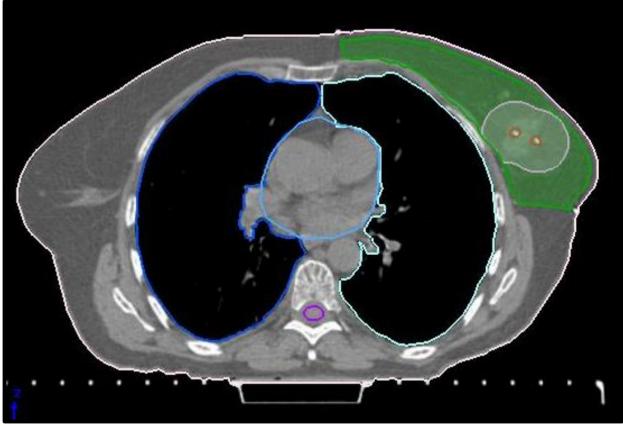


EBRT allows for conformal treatment

• In pre operative or post operative (several ongoing trials)

• Positioning and contouring are essential: more risks to miss the target!

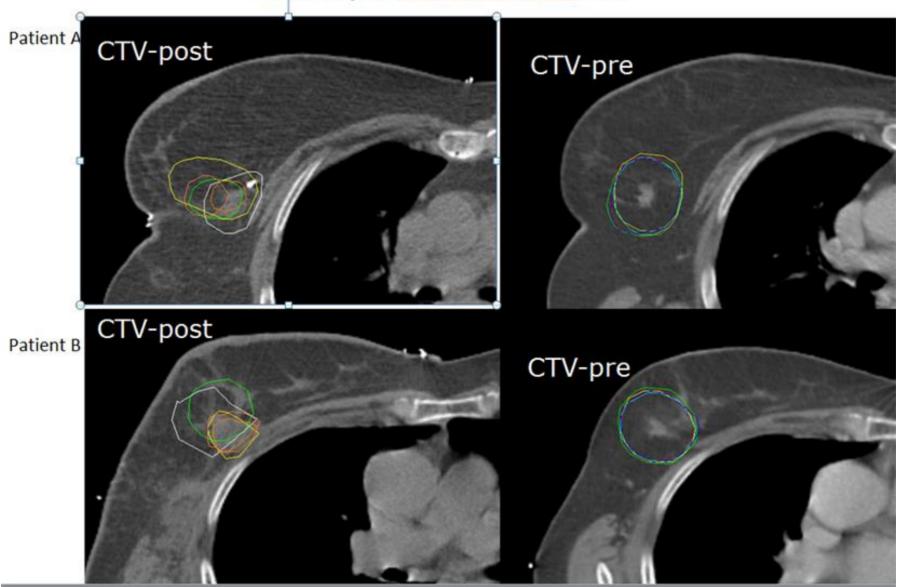






Preop. vs postop. delineation

van der Leij et al Radiother Oncol 2014





PAPBI: first résults

Before RT

6 months

12 months

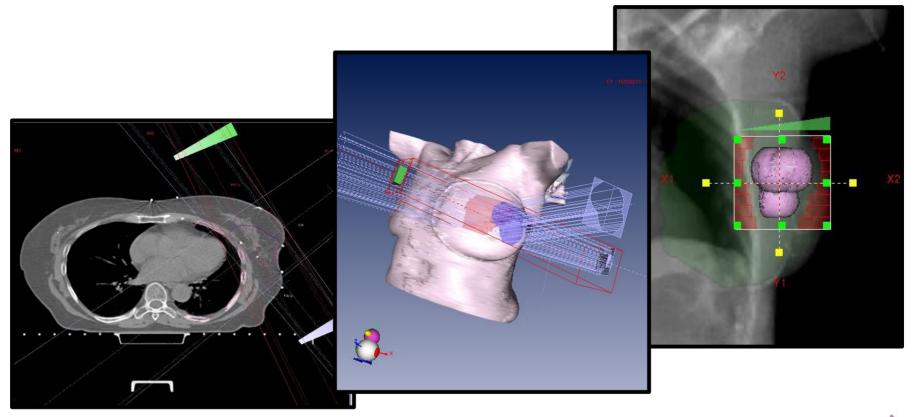
24 months



Dosimetry

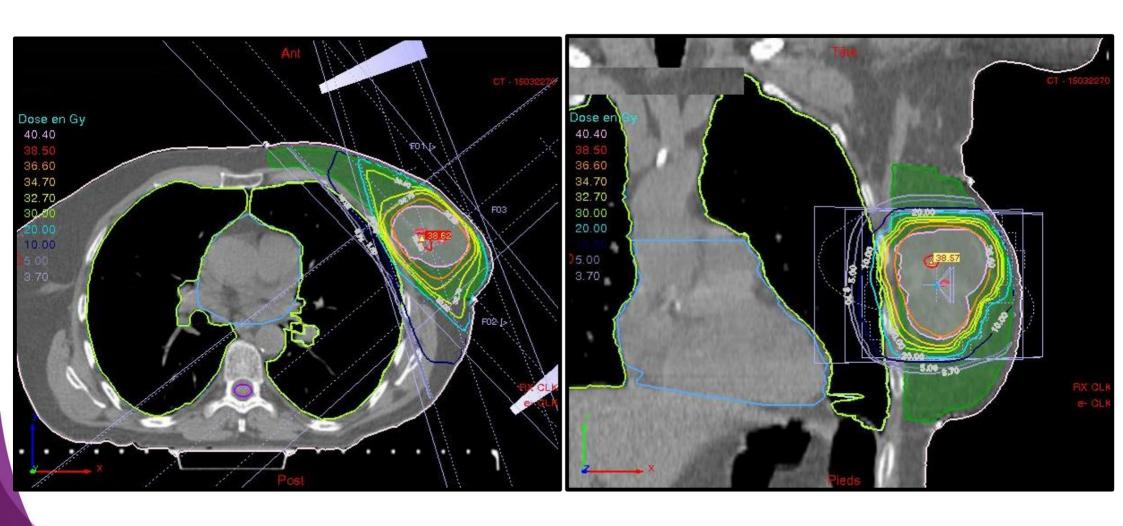
Technique used in routine at Gustave Roussy:

- 2 tangential 6MV photon beam
- 1 direct electron beam





Dosimetry

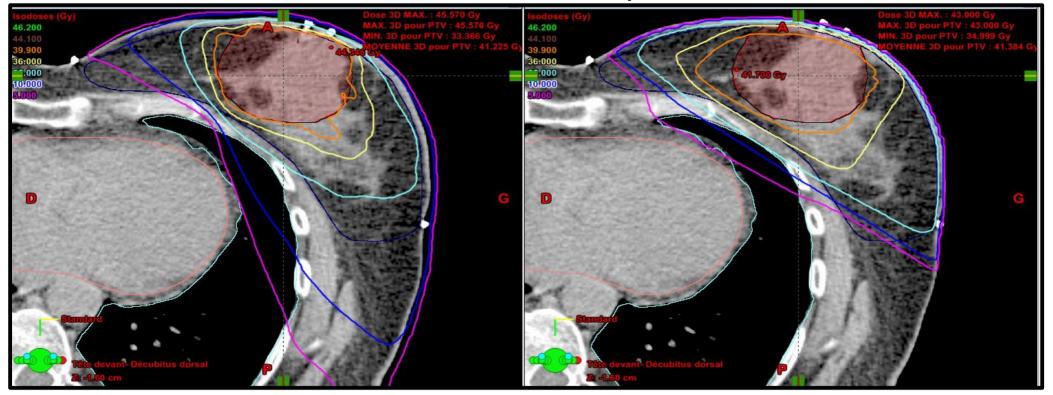




Dosimetric comparision between APBI

Rapidarc technique

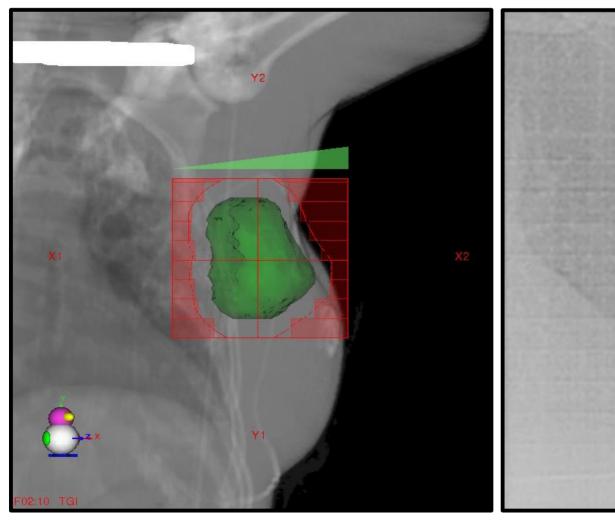
3D conformal with photons and electrons

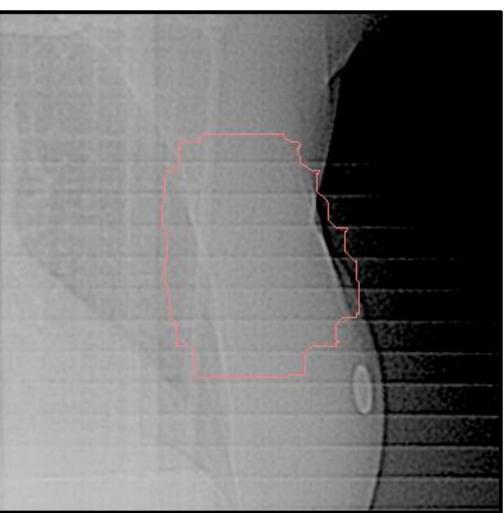


Advantages	Drawbacks
Lower heart dose	Higher hot spot
Lower whole breast dose	Increased low doses to lung



Re-positioning control

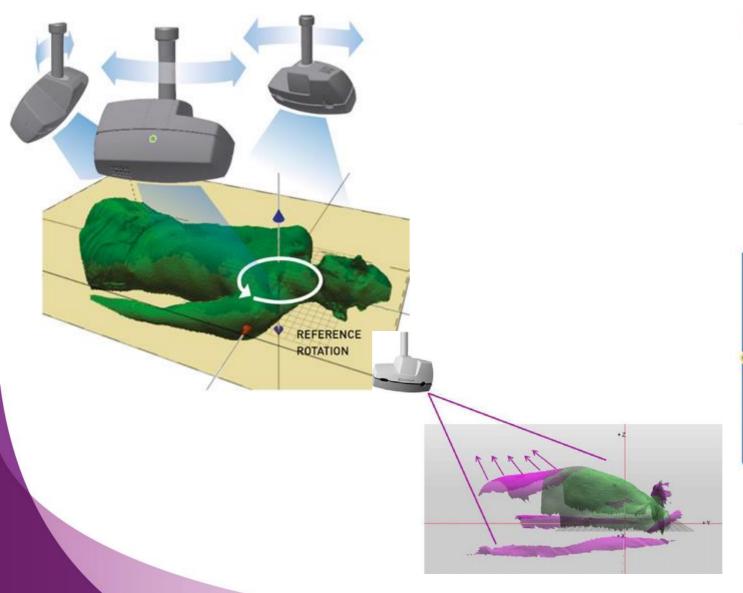


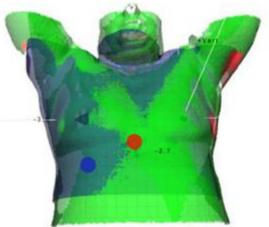


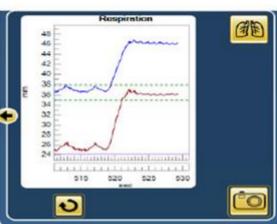


Positioning control

• Ex: C-RAD Catalyst system



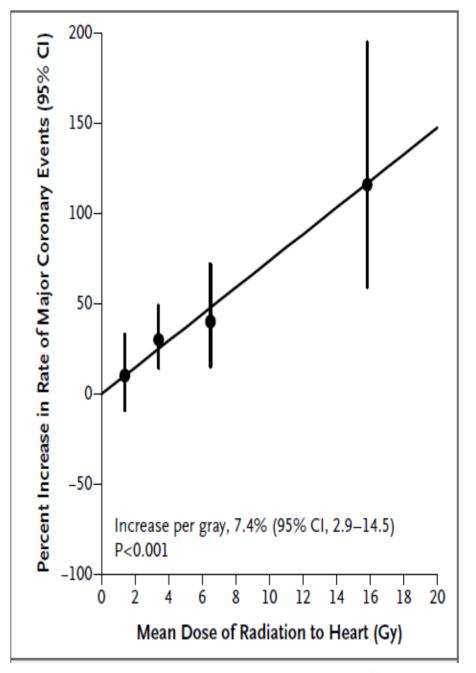






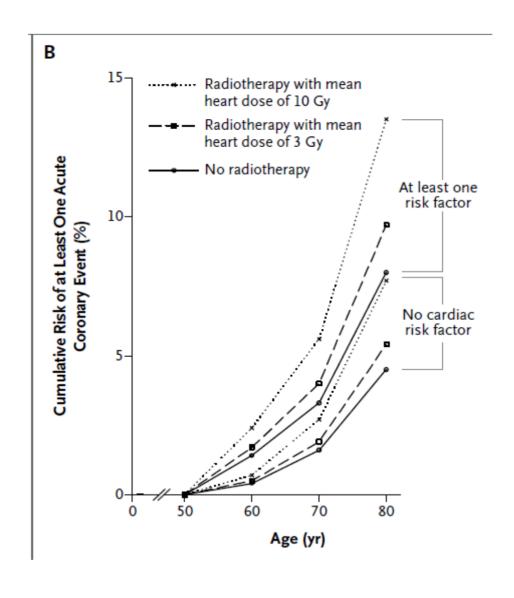
Heart Toxicity

"The overall average of the mean doses to the whole heart was 4.9 Gy (range, 0.03 to 27.72). Rates of major coronary events increased linearly with the mean dose to the heart by 7.4% per gray (95% confidence interval, 2.9 to 14.5; P<0.001), with no apparent threshold. The increase started within the first 5 years after radiotherapy and continued into the third decade after radiotherapy. The proportional increase in the rate of major coronary events per gray was similar in women with and women without cardiac risk factors at the time of radiotherapy"

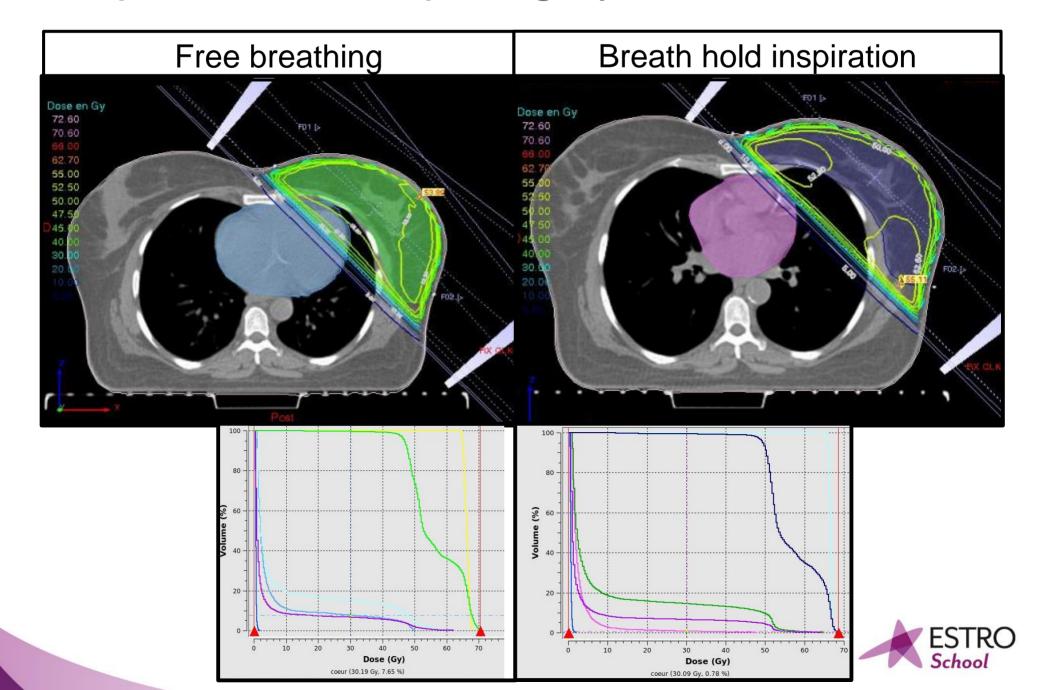




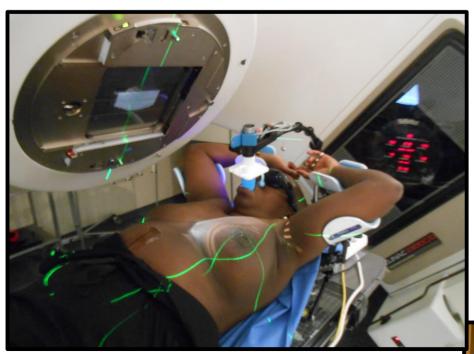
Cardiac risk is increased by cardiovascular risk factors



Improved heart sparing by breath hold



How to improve heart sparing?



• Inspiration breath hold technique









- Accelerated hypofractionated whole breast and partial breast irradiation are changing our practices for early breast cancers with good prognosis factors
- Contouring and positioning remain key points for these treatment strategies
- Moving toward better sparing OAR means we need to assess low dose consequences as well





Breast case – Physics or metaphysics?

Peter Remeijer

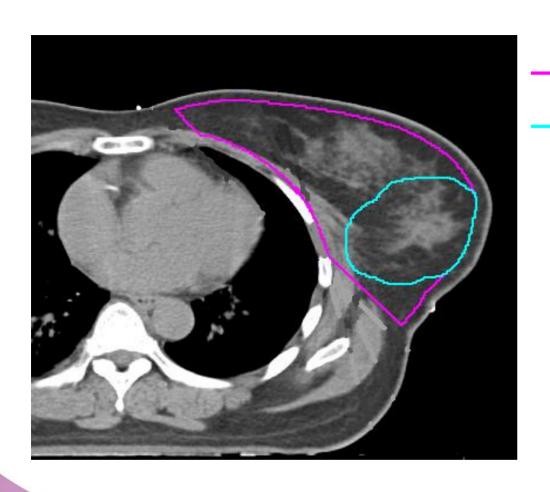
Department of Radiation Oncology

The Netherlands Cancer Institute





Common target volumes

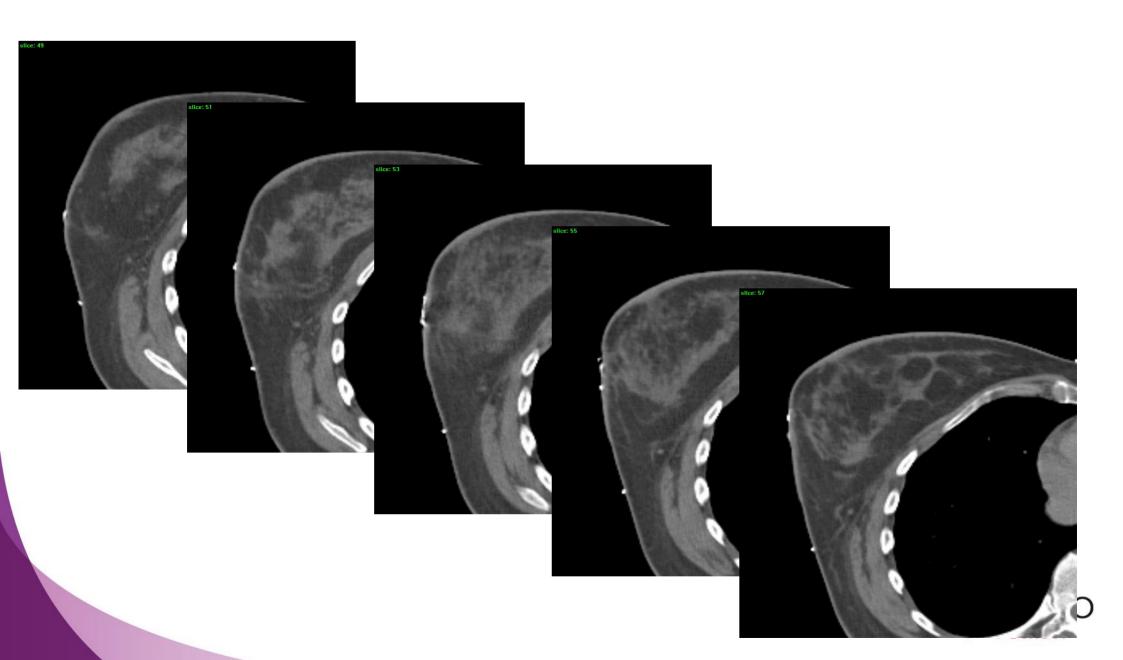


Whole breast (50 Gy)

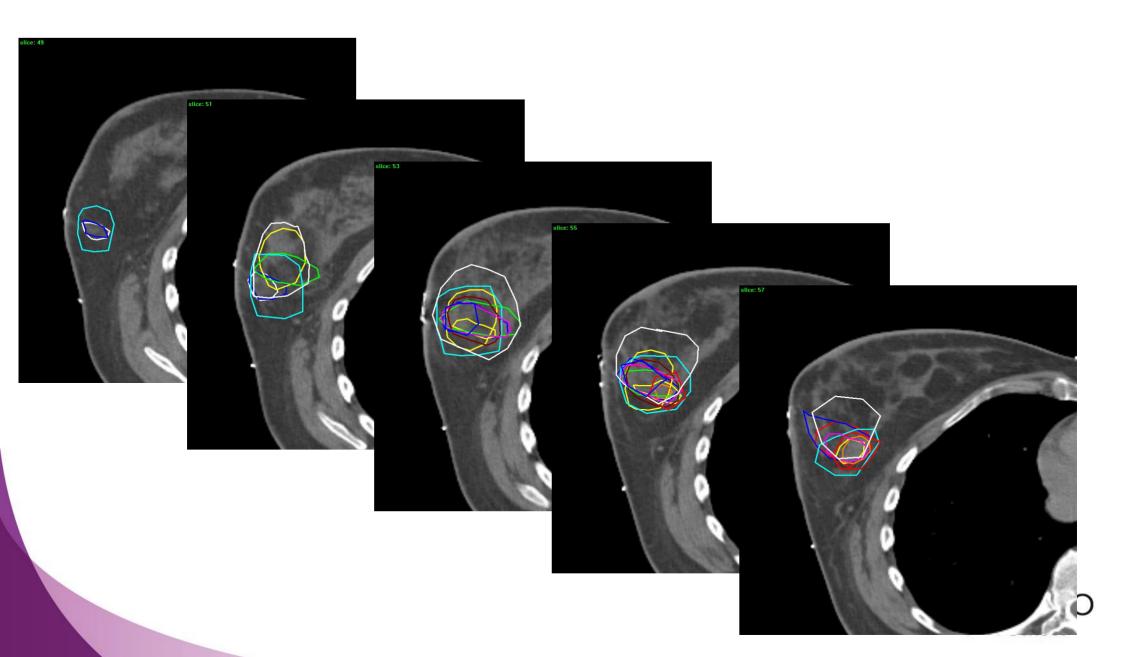
Excision cavity (16 Gy)



Target volume delineation - variability



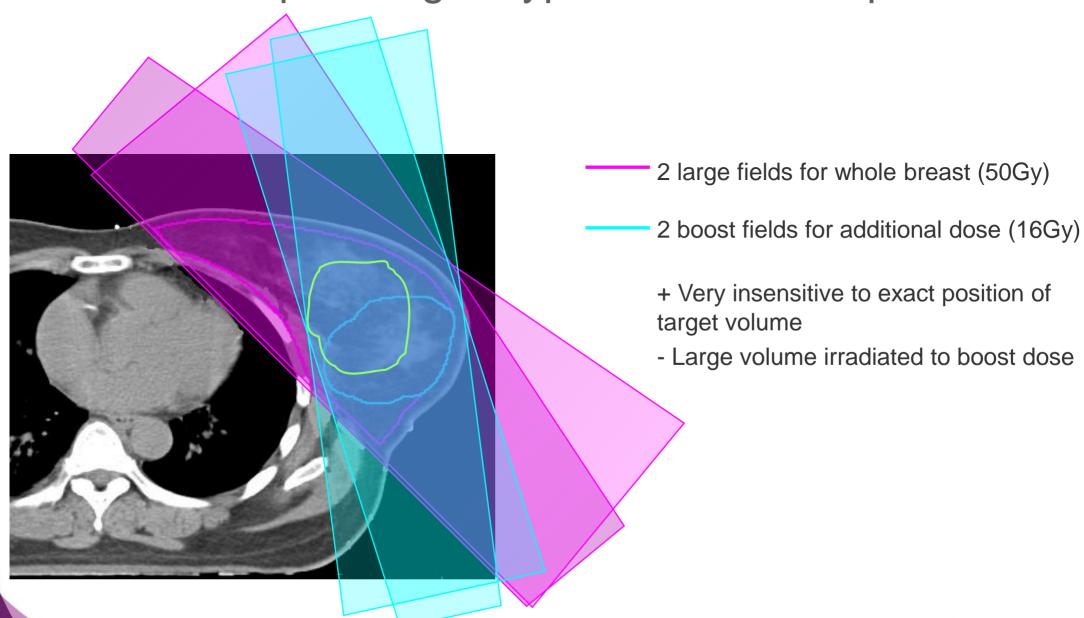
Target volume delineation - variability

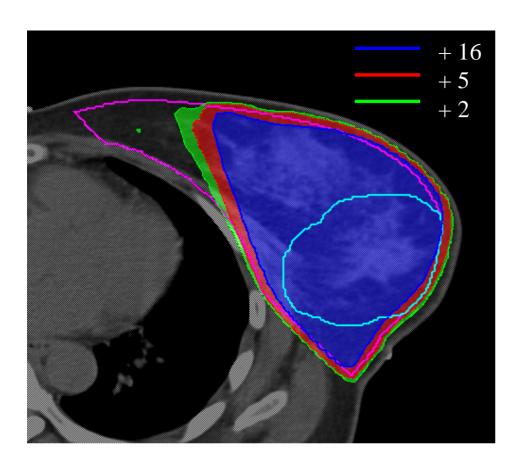


Target volume delineation - variability

- Possible causes
 - Different opinion of the clinicians
 - Image quality
- Possible solutions
 - Clear protocols, good collaboration between OR,
 Pathology, RT
 - Markers
 - Registration of pre-and post-op imaging (difficult!)
 - Multiple modalities

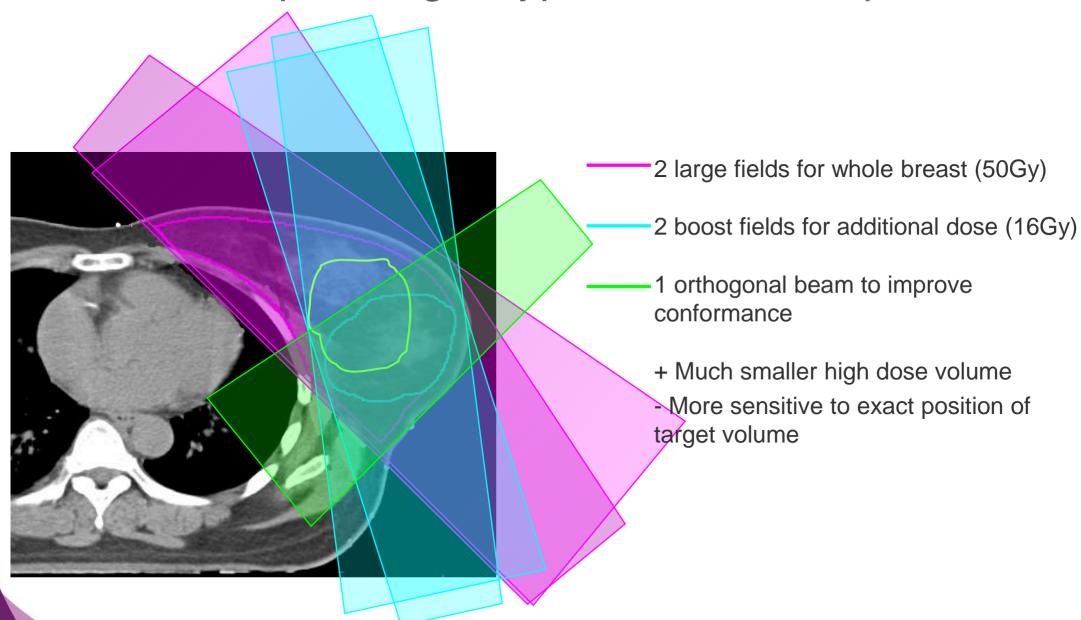


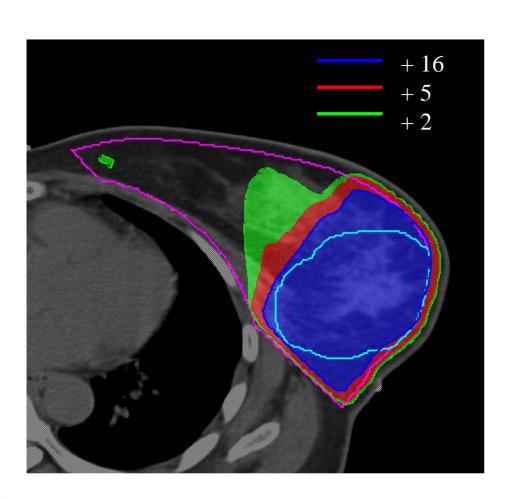




- + Very insensitive to exact position of target volume
- Large volume irradiated to boost dose







- + Much smaller high dose volume
- Sensitive to exact position of target volume
- → Image guidance / position verification



Changes during treatment



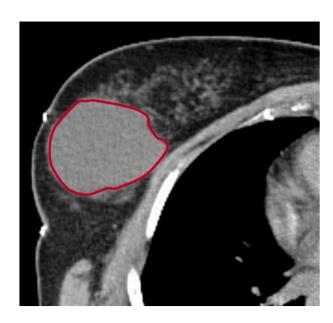
Setup errors

	No-correction _{meas}				Offline _{meas}				
		CBCT				CBCT			
		CC	LR	AP		CC	LR	AP	
Σ (mm)		3.8	3.1	2.5	Ī	1.7	1.4	1.2	
σ (mm)		2.8	2.2	2.6		3.1	2.3	3.0	
2.5Σ + 0.7σ (mm)		11.5	9.2	8.0		6.4	5.2	5.1	



Seroma changes

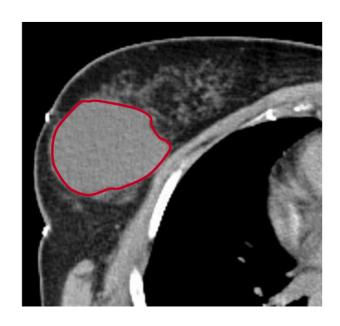
- Possibly occurs after breast sparing surgery
- Fluid in excision cavity



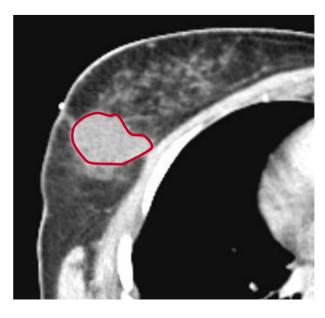
Boost is generally based on this seroma volume

Seroma changes

Seroma shrinkage



Planning CT

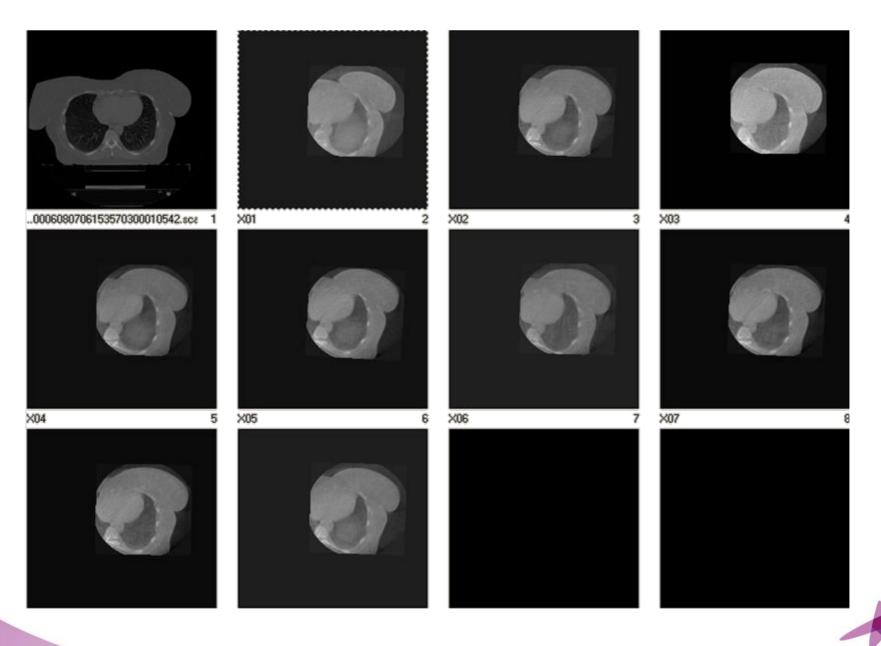


Repeat CT (during treatment)

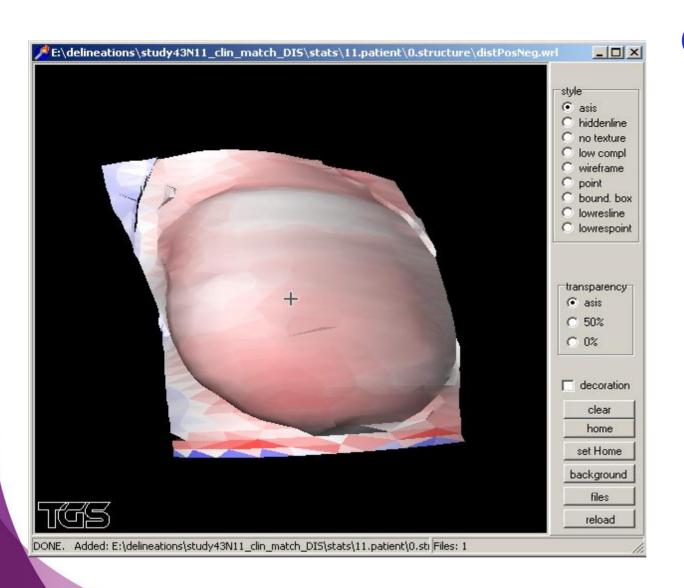
Can be partially solved by adaptive RT

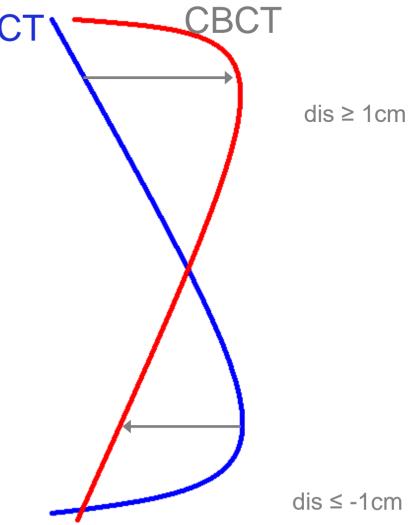


Changes in breast shape

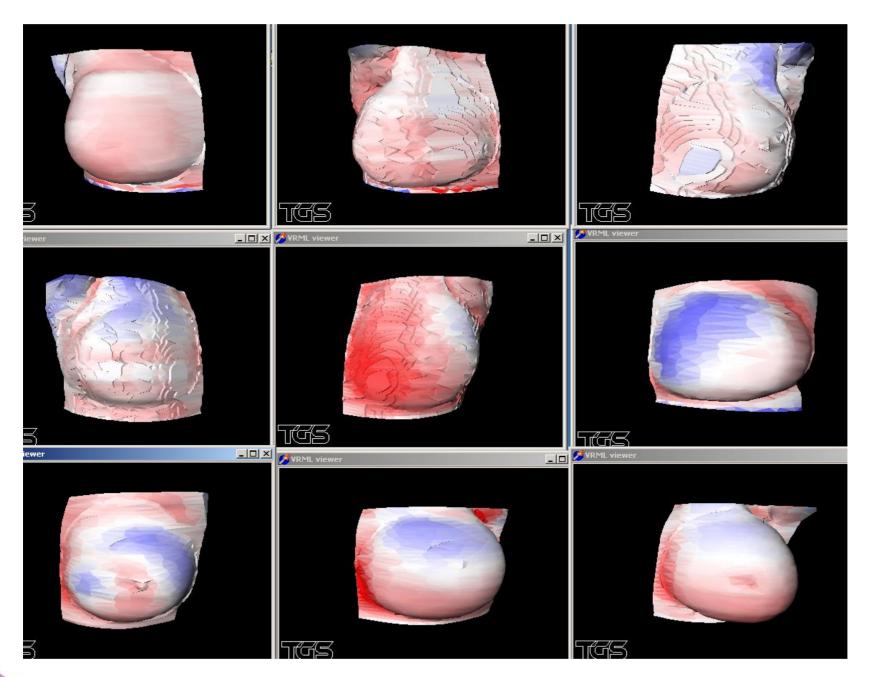


Average difference (treatment → planning)



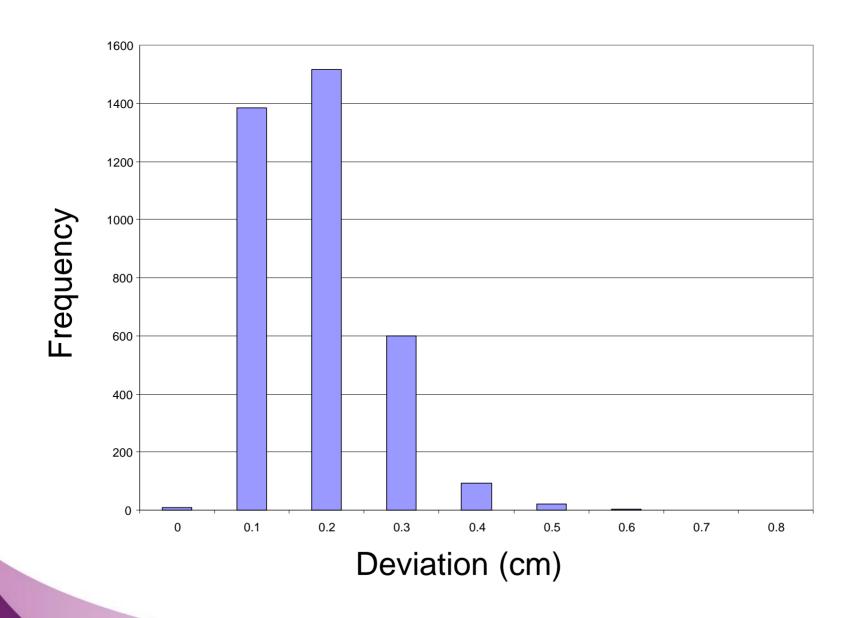








Average difference (treatment → planning)





Margins



- Clinically used margin for breast: o mm!
 - Adapted from sim-technique it's just beam setup
 - Clearly not enough according to conventional margin ideas
- Clinically used margin for boost: 5 mm
 - Let's see if that's enough

	Systematic	Random
Delineation	2.0 mm	_
Setup	1.5 mm	2.5 mm
Shape changes	2.0 mm	2.0 mm
Total	3.2 mm	3.2 mm

- Margin: 2.5 * 3.2 + 0.3 * 3.2 = 9 mm



• So why is this not leading to lots of local recurrences?



- For the whole breast
 - It's a CTV. Small underdosage does not necessarily underdose actual tumor cells
 - → Risk of tumor cell underdosage small
 - Ongoing debate whether it's even necessary to treat the whole breast



- For the boost
 - It's a boost with a 50 Gy background dose, so severe underdosage will not occur
 - CTV margin for the excision cavity is usually large → compensates for small PTV margin
 - Conformity is not very good with current planning techniques → effectively the margin is bigger
 - This is however **not** the case for partial breast treatments



Take home messages

• Conventional treatment techniques are not very critical with respect to geometrical uncertainties

• Partial breast treatments will be more critical because of lack of background dose and more advanced and conformal treatment techniques (e.g. VMAT)





Breast IGRT: An RTT Perspective

Liz Forde, RTT
Assistant Professor
The Discipline of Radiation Therapy
School of Medicine
Trinity College Dublin







Fundamental IGRT Questions

- When should I image?
 - > Frequency
- **How** should I image?
 - > Technology
 - Projection
- What can I see?
 - > What is my target
- What should I match to?
 - > Surrogate for target position





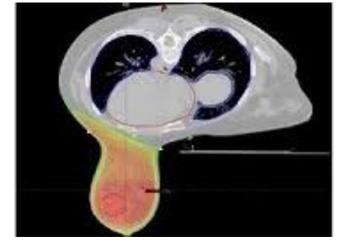
Site Specific Points to Consider

- Laterality
 - Right/Left
 - Cardiac dose
- Patient positioning
 - Supine, Prone or lateral decubitus
- Target volume
 - Whole or Partial Breast
 - Boost
- Simulation
 - > 3D or 4D
- Breathing motion
 - DIBH
 - Free breathing

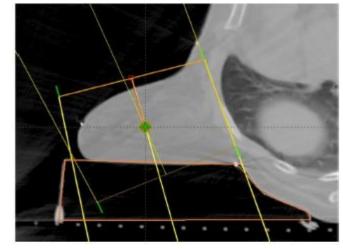


All of these factors will influence how we image this patient group

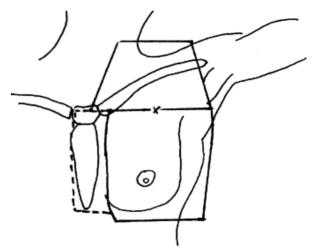




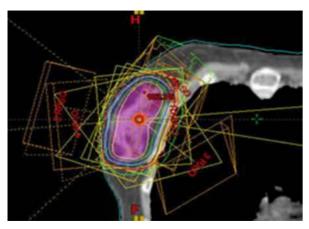
Prone



Lateral decubitus



Supine: IMC (ph/e junx)



APBI



Electron boost to surgical bed







DIBH





On Treatment Verification

- · Look! There is it! I can see the target!
 - Whole breast RT
- Confirm gross external positioning information
 - Light field
 - > FSDs
- What else do we want to see?
 - Contour changes
 - > Tumour bed
 - > Seroma
 - Surgical clips





Match Anatomy

- Breast contour
- Lung volume
- Ribs
- Seroma
- Surgical Clips



Surgical Clips

- Act as a surrogate for the tumour bed
- Improve accuracy in delineation and used for positional verification
- Clip insertion after breast conservation surgery
- Caution artefact on planning CT
 - Impact on electron beam dosimetry?
- Either use directly in match or export isodose lines from planning to ensure they fall within required dose
 - Donovan et al., 2012
 - Similar to Post Prostatectomy clips



On Treatment IGRT

- Largely driven by what is available to you
- Make the most of it
- Consider the clinical impact
 - > Tighter margins?

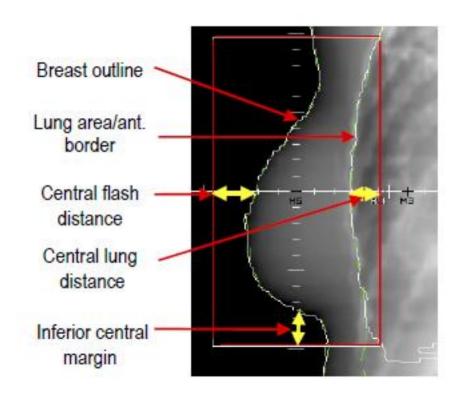


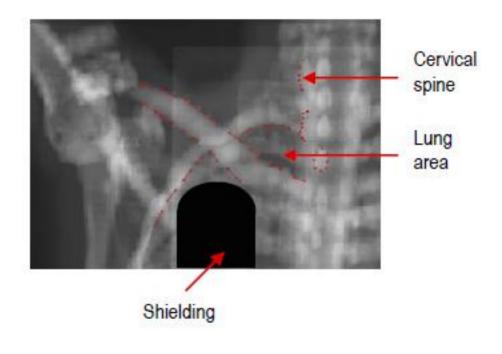
MV_{2D}

- Widely available
- Ability to acquire continuous "snapshot" during the fraction
- Will provide field border information
- Will provide assessment of lung volume, breast contour
- Adequate for whole breast RT with standard fractionation
- Typically 5mm tolerance is acceptable
- Difficult to visualise surgical clips
- Depending on lung in field, generally sufficient information from a "single" acquisition



MV_{2D}







2D/2D (Paired orthogonal 2D)

Used for isocentre position check

Field border information is not displayed

A minimum requirement for all advanced techniques

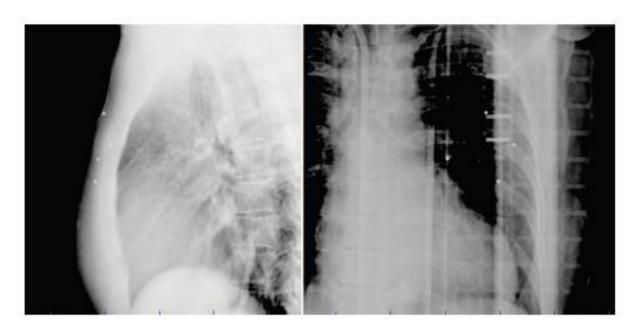


FIG. 2. Anterior–posterior and lateral paired kV images of a patient on treatment day 1. Yue at al., 2013

kV decreases dose burden and increases image quality



3D (CBCT)

Provides:

- > Isocentre position verification
- Internal soft tissue anatomy
- Clearer image of clips
- > Information on changes in target during treatment
 - Seroma changes

Consider:

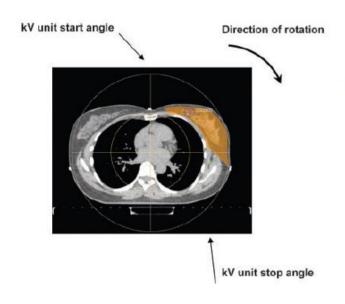
- Dose
- Collision risk
- Ease of accurate registration and match



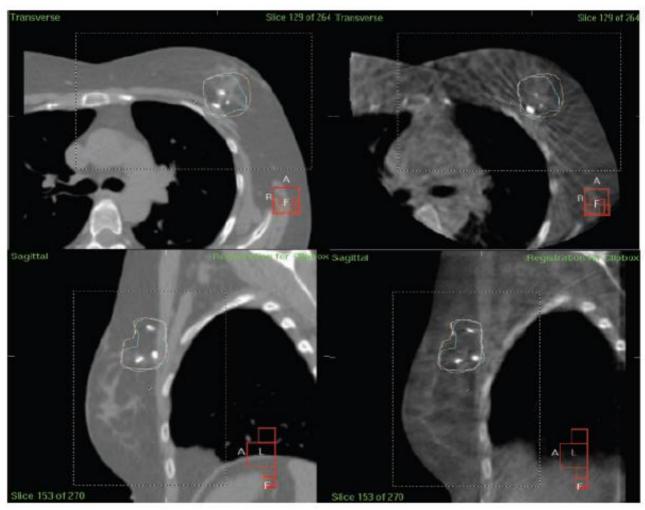
3D (CBCT)

- Limitations
 - Collision
 - > Field of view
 - Increased dose to contra lateral breast
- CBCT not acquired at the isocentre to avoid collision
- Then once matched the shift includes the offset from isocentre position
- Adds time and potential errors
- Donovan et al. (2012) stipulate limitations on iso position to account for this

3D (CBCT): Clarity of Surgical Clips



Isodose lines have been exported to confirm coverage







3D (CBCT): Clarity of Surgical Bed

Setup error for EPID and cone-beam CT ● R. TOPOLNJAK et al.

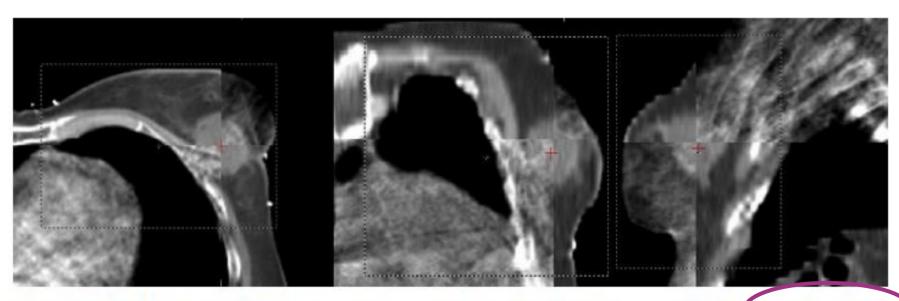


Fig. 2. Match of planning computed tomography (CT) and cone-beam CT (CBCT) images after bory (stemum and rib) anatomy registration. A bony anatomy rigid registration was performed based on image data in a user defined, three-dimensional, box-shaped region of interest (white dashed line) using a chamfer matching algorithm.

Topolnjak et al., 2009



Video-Based Surface Mapping

- Whole surface shape matching
- Provides surface anatomy information and can demonstrate the impact of breathing
- Can this be correlated to provide shift/positional information?
 - Often used in conjunction with other imaging devices
- No additional radiation







A Look at the Literature

Table 4.5 Articles that discuss doses, anatomy matching methods and seroma visualisation

KV imaging Method	Author	Matched Method	Sample Size>20	Seroma Visible	Surgical clips	Auto co- registration	Reported imaging dose to patient.	Safe to acquire at iso centre
CBCT	Jain et al.(2009)	Bones	No	No	No	Yes	Yes	No
	White et al. (2007)	Lung/external contour	Yes	Yes	No	Potential	Yes	Yes
	Kim et al. (2007)	Clips	No	Yes	Yes	Yes	Yes	No
	Topolnjak et al. (2010)	Sternum/ribs	Yes	Unknown	No	Yes	No	Unknown
	Yang et al.(2010)	Unknown	No	Yes	No	Yes	Yes	Unknown
	Donovan et al.(2012)	Clips	Yes	Yes	Yes	Yes	Yes	Yes
kV*	Yue et al. (2011)	Bony to gold fiducials	Yes	Yes	Yes	Unknown	No	Unknown
	Lawson et al. (2008)	Bony	Yes	Unknown	Yes	No	Yes	Unknown
kV* vs. CBCT	Fatunase et al. (2008)	Bones, then soft tissue	No	Yes	Yes	No	Yes	No for both.

*orthogonal kV imaging

L. Lewis Improving Radiotherapy for Breast Cancer: Identification of the tumour bed and characterisation of target volume changes. 2013 MSc Thesis, available online



What is Happening in Europe?

• 2010 Survey of EORTC affiliated institutions

• "Electronic portal imaging for patient set-up is used by **92%** of the institutions." (van der Laan et al., 2010)

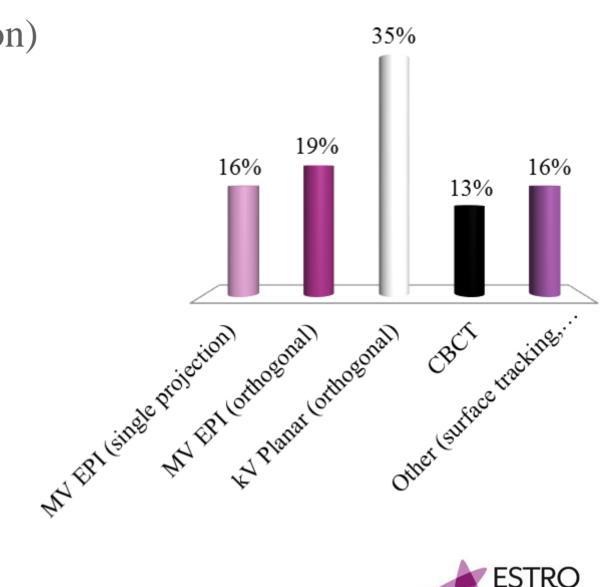
So what does Europe look like in 2017?

What is happening in Europe today?



In my clinical department, for standard WBRT, we image using:

- A. MV EPI (single projection)
- B. MV EPI (orthogonal)
- C. kV Planar (orthogonal)
- D. CBCT
- E. Other (surface tracking, Tomotherpy)





How did you compare with The US? 2016 Survey of ASTRO Members (Nabavizadeh et al., 2016)

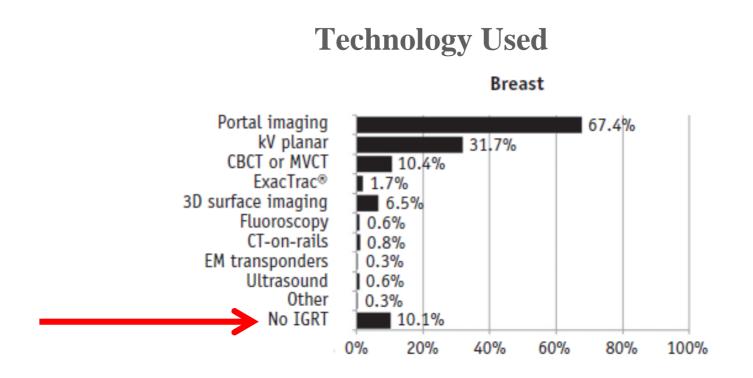
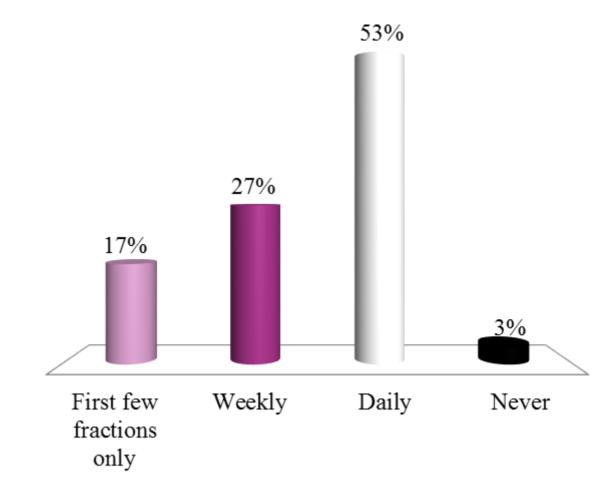


Fig. 2. Physician-reported image guided radiation therapy frequency (black) and on-line image verification frequency (gray) for standard fractionation treatments, by disease site. *Abbreviations:* 3D-CRT = 3-dimensional conformal radiation therapy; fx = fractions; IMRT = intensity modulated RT.



In my clinical department, for standard WBRT, we image:

- A. First few fractions only
- B. Weekly
- C. Daily
- D. Never





How did you compare with The US? 2016 Survey of ASTRO Members (Nabavizadeh et al., 2016)

Frequency of Imaging

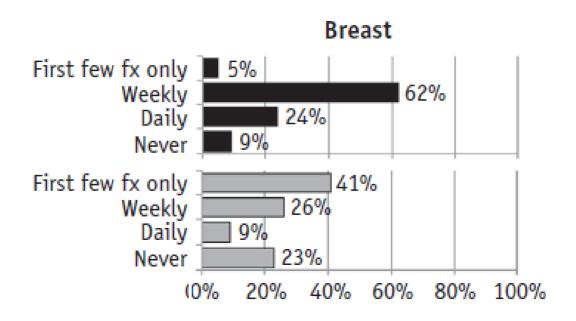


Fig. 2. Physician-reported image guided radiation therapy frequency (black) and on-line image verification frequency (gray) for standard fractionation treatments, by disease site. *Abbreviations:* 3D-CRT = 3-dimensional conformal radiation therapy; fx = fractions; IMRT = intensity modulated RT.



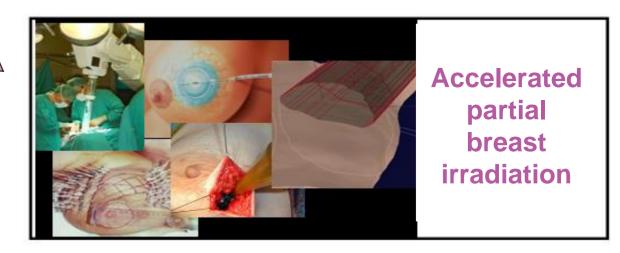
Therapeutic strategy: Which radiotherapy?

Two changing practice concepts have modified the standard whole breast irradiation 50Gy +/- boost has been replaced





Whelan NEJM 2010; START A and B Lancet Oncol 2008



Vaidva Lancet 2010; Bourgier IJROBP 2010; Lemanski IJROBP 2010; Taghian IJROBP 2005; Polgar IJROBP 2004; Vicini IJROBP 2003; Formenti IJROBP 2003;



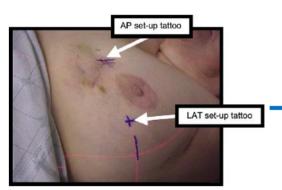


Fig. 1. Patient with skin marker crosshairs representing isocentric setup at surgical cavity.

IGRT for (Supine) APBI: What are people doing?

Int. J. Radiation Oncology Biol. Phys., Vol. 76, No. 2, pp. 528–534, 2010

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0360-3016/10/\$-see front matter

doi:10.1016/j.ijrobp.2009.02.001

One of the first reports on IGRT APBI

CLINICAL INVESTIGATION

Breast

CLINICAL EXPERIENCE WITH IMAGE-GUIDED RADIOTHERAPY IN AN ACCELERATED PARTIAL BREAST INTENSITY-MODULATED RADIOTHERAPY PROTOCOL

CHARLES E. LEONARD, M.D.,* MICHAEL TALLHAMER, M.S.,* TIM JOHNSON, Ph.D.,*
KARI HUNTER, C.M.D.,* KATHRYN HOWELL, M.D.,* JANE KERCHER, M.D.,† JODI WIDENER, M.D.,†
TERESE KASKE, M.D.,‡ DEVCHAND PAUL, M.D.,§ SCOT SEDLACEK, M.D.,§ AND DENNIS L. CARTER, M.D.,*

*Rocky Mountain Cancer Centers, Littleton, CO; †Arapahoe Surgical Associates, Littleton, CO; †Sally Jobe Diagnostic Breast Center, Greenwood Village, CO; and §Rocky Mountain Cancer Centers, Rose Hospital, Denver, CO

Orthogonal **MV** images taken **daily**

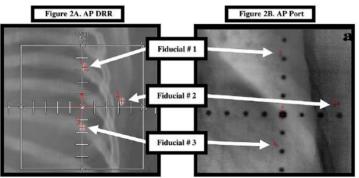


Fig. 2. Anterior/posterior digital reconstruction radiograph and corresponding port film (arrows indicate respective fiducial

Imaging dose included in plan

Table 5. Individual and total dose contribution of AP and lateral port films for 5 patients over the course of accelerated partial breast radiotherapy to the isocenter and 100% of PTV using image-guided radiotherapy

	ISO dose (cGy)					
Patient	AP	Lateral	Total	100% PTV		
1	30.7	25.3	56	45		
2	28.1	30.7	58.8	48		
3	26.4	30.1	56.5	50		
4	29.7	28	57.6	49		
5	30.3	29.9	60.2	54		
As a percentage of prescription dose of 3,850 cGy						
1	0.8	0.7	1.5	1.2		
2	0.7	0.8	1.5	1.3		
3	0.7	0.8	1.6	1.3		
4	0.8	0.7	1.5	1.3		
5	0.8	0.8	1.6	1.4		

IGRT for (Supine) APBI: What are people doing?



Int. J. Radiation Oncology Biol. Phys., Vol. 76, No. 2, pp. 528–534, 2010

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0360-3016/10/8–see front matter

doi:10.1016/j.ijrobp.2009.02.001

CLINICAL INVESTIGATION

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CHARLES E. LEONARD, M.D.,* MICHAEL TALLHAMER, M.S.,* TIM JOHNSON, Ph.D.,* KARI HUNTER, C.M.D.,* KATHRYN HOWELL, M.D.,* JANE KERCHER, M.D.,† JODI WIDENER, M.D.,† TERESE KASKE, M.D.,‡ DEVCHAND PAUL, M.D.,§ SCOT SEDLACEK, M.D.,§ AND DENNIS L. CARTER, M.D.*

*Rocky Mountain Cancer Centers, Littleton, CO; †Arapahoe Surgical Associates, Littleton, CO; †Sally Jobe Diagnostic Breast Center, Greenwood Village, CO; and §Rocky Mountain Cancer Centers, Rose Hospital, Denver, CO

Because of the reliability of these fiducial markers, we have reduced the size of port films for IGRT. Before the use of fiducials, reviewing physicians required visualization of the surrounding anatomy, specifically the ribs, clavicle, and sternum. This required the more traditional double-exposure port with the second larger field exposure. When it is not necessary to view surrounding anatomical landmarks because of fiducial placement in proximity to the target cavity, it is possible to reduce port field sizes.

cerning their use in partial breast treatment. This could suggest that margins might be reduced for a smaller PTV volume than is used currently. Up to this time, an additional margin of 1 cm from the CTV had been used. However, owing to the use of these fiducial markers, this additional margin may be reduced by 5 mm. This would be well within two standard deviations of the average mean error of our IGRT experience.

IGRT for (Supine) APBI: What are people doing?

Distinction b/w surgical clips and fiducials

Textured gold fiducials for stability and visualisation

Daily orthogonal MV EPI

Clinical Investigation. Breast Cancer

Published 2012

Validating Fiducial Markers for Image-Guided Radiation Therapy for Accelerated Partial Breast Irradiation in Early-Stage Breast Cancer

Catherine K. Park, M.D., M.P.H.,* Jakub Pritz, M.S.,† Geoffrey G. Zhang, Ph.D.,* Kenneth M. Forster, Ph.D.,* and Eleanor E.R. Harris, M.D.*

From the *Department of Radiation Oncology, H. Lee Moffitt Cancer Center & Research Institute, Tampa, FL; and †Department of Physics, University of South Florida, Tampa, FL

Received Jan 12, 2011, and in revised form Jun 22, 2011. Accepted for publication Jul 18, 2011

Visualisation of fiducials on 100% MV images Centre of fiducials correlated to centre of seroma When matching to fiducials margins can be reduced to 6mm compared to bone (10mm)



Aim: to assess the residual and intrafraction errors

IGRT for (Supine) APBI: What are people doing?

PTV = CTV+10mm 5 fld non coplanar 95%/95%

Cai et al. Radiation Oncology 2010, 5:96 http://www.ro-journal.com/content/5/1/96



RESEARCH Open Access

Impact of residual and intrafractional errors on strategy of correction for image-guided accelerated partial breast irradiation

Gang Cai^{1†}, Wei-Gang Hu^{1*†}, Jia-Yi Chen^{1*}, Xiao-Li Yu¹, Zi-Qiang Pan¹, Zhao-Zhi Yang¹, Xiao-Mao Guo¹, Zhi-Min Shao², Guo-Liang Jiang¹

Pre and post fx XVI
Grey value match
Manual adjustment with:
skin, chest wall and clips
2-3 mins
Matched by RO

CBCT does not guarantee absolute accuracy 13mm margin required to account for initial setup and intrafraction errors



MRI Based IGRT – The Future?

Clinical Investigation

Magnetic Resonance Image Guided Radiation Therapy for External Beam Accelerated Partial-Breast Irradiation: Evaluation of Delivered Dose and Intrafractional Cavity Motion

Sahaja Acharya, MD, Benjamin W. Fischer-Valuck, MD, Thomas R. Mazur, PhD, Austen Curcuru, BS, Karl Sona, MS, Rojano Kashani, PhD, Olga Green, PhD, Laura Ochoa, ANP, PhD, Sasa Mutic, PhD, Imran Zoberi, MD, H. Harold Li, PhD, and Maria A. Thomas, MD, PhD

Department of Radiation Oncology, Washington University School of Medicine, St. Louis, Missour

Received Jan 9, 2016, and in revised form Jul 8, 2016. Accepted for publication Aug 10, 2016.



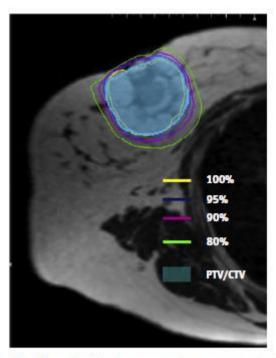


Fig. 1. Plan visualized on magnetic resonance simulation. Planning target volume (PTV) (= clinical target volume [CTV]) is shown in light blue color wash. Isodose lines: 100% (yellow), 95% (dark blue), 90% (magenta), 80% (green). (A color version of this figure is available at www.redjournal.org.)

Take Home Message

- There is an abundance of imaging technologies and strategies available for this site
- IGRT for breast is largely dependant not only what is available to you, but the planning technique that is used
- Advanced treatment techniques require more sophisticated imaging techniques
 - > APBI, IMRT, VMAT



Lung

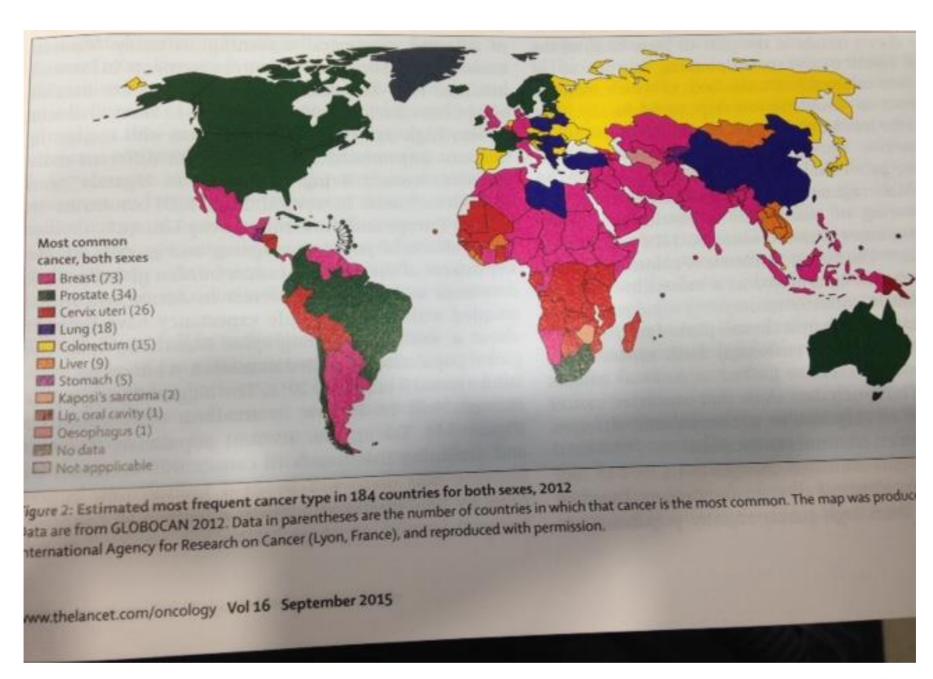




Jose Lopez, M.D., Ph.D Radiation Oncology University Hospital Virgen del Rocio Seville, Spain

Advanced skills in modern radiotherapy

Prague, Czech Republic –11-15 June 2016





Outline of Talk

• Preclinical rationale behind **oligometastatic** state

• Clinical data supporting benefit to local treatment in oligometastatic NSCLC

Case report

• Discussion of current multidisciplinary (physician, phisyc and RTTs) management

Introduction

- Definitive radiotherapy has historically been reserved for patients with stage I-III disease.
- The most common indication for RT in patients with metastatic lung cancer has been palliation for pain or other symptoms
- However, <u>stage IV lung cancer is a very broad category</u>, and prior studies have suggested that some patients with stage IV lung cancer and only a few distant metastases
 ('oligometastasis') may benefit from local therapy to both the primary tumor and the distant sites of disease.



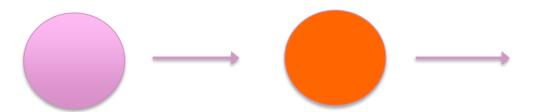
Introduction

- Spectrum of metastatic patients exists
 - > Indolent vs. aggressive course
- In-between locoregionally confined and true metastatic state, there appears to exist intermediate state of low disease burden systemically=oligometastasis
 - Can these patients be "cured"?



"Oligo" means "having few, having little."

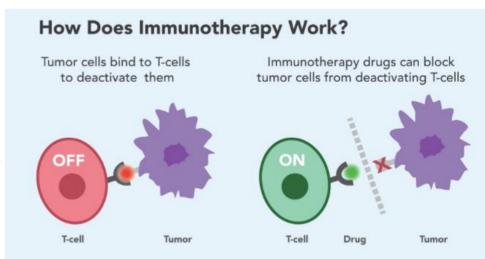
Studies with lung cancer have defined oligometastatic disease as up to 5 metastatic lesions.



"Oligo" means "having few, having little." **Studies with lung cancer have** defined oligometastatic disease as up to 5 metastatic lesions. Adrenal gland Dr Daniel Gomez. MD Anderson Cancer Center

Recent Trials Addressing Management of Oligometastatic NSCLC

- Recent developments
 - > Targeted agents
 - Maintenance chemotherapy
 - ➤ Technologic advances permitting ablative doses of radiation therapy



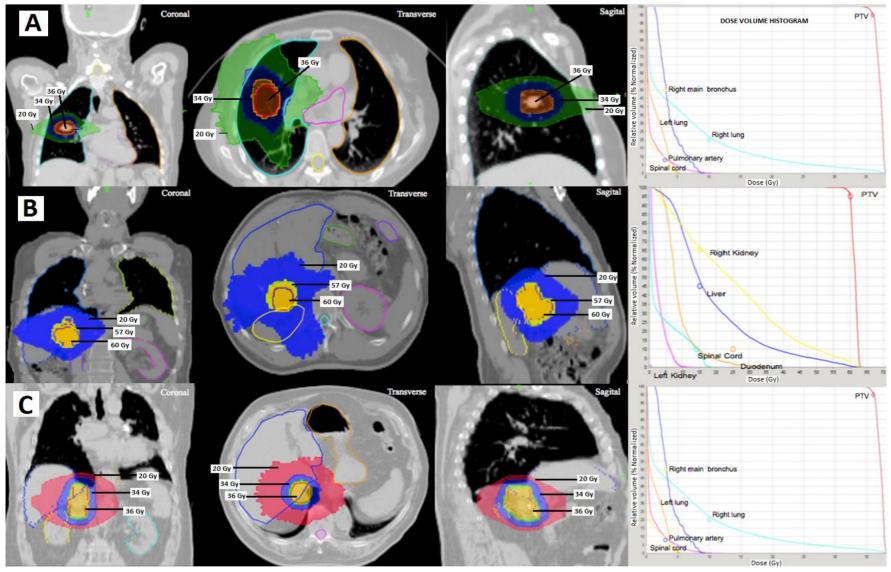


Clinical Data Supporting Local Treatment in Oligometastatic Setting

Author	Year	n	Timespan (years)	Single institution
Twomey	1982	2	14	Yes
Reyes	1990	5	4	Yes
Raviv	1990	3	nr	Yes
Kirsch	1993	1	6	Yes
Higashiyama	1994	5	12	Yes
Ayabe	1995	3	9	Yes
Urschel	1997	1	9	Yes
Bendinelli	1998	4	4	Yes
Tsuji	1998	1	2	Yes
Linos	1998	1	1	Yes
Porte	1998	11	8	Yes
Wade	1998	14	7	No (159 centers)
de Perrot	1999	1	5	Yes
Bretcha-Boix	2000	5	nr	Yes
Ambrogi	2000	5	7	Yes
Porte	2001	43	12	No (8 centers)
Mercier	2004	23	14	Yes
Lucchi	2005	11	10	Yes
Pfannschmidt	2005	11	7	Yes
Sebag	2006	9	9	Yes
Munoz	2006	1	5	Yes
Strong	2007	29	11	Yes



STEREOTACTIC ABLATIVE RADIOTHERAPY DELIVERED BY HELICAL TOMOTHERAPY FOR EXTRACRANEAL OLIGOMETASTASIS





STEREOTACTIC ABLATIVE RADIOTHERAPY DELIVERED BY HELICAL TOMOTHERAPY FOR EXTRACRANEAL OLIGOMETASTASIS

CONTOURS

- GTV: defined only as the solid abnormality on CT + PET
- ITV: using a multiple CT scan (free breathing, maximal inspiration, and maximal expiration)
- PTV: 0.5 cm in the axial plane and 1.0 cm in the craneocaudal plane

DOSE PRESCRIPTION

- Lung (not chest wall): 3 fractions of 20 Gy
- Lung (chest wall): 3-5 fractions of 12 Gy for lesions
- Lung (central): 8 fractions of 7.5 Gy

CHEMOTHERAPY (90%)

- FOLFOX/FOLFIRI

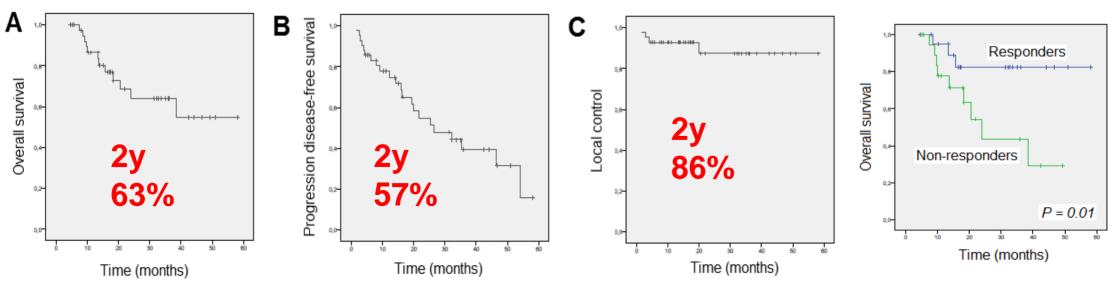


DOSE CONSTRAINTS

THORAX

- Chronic lung disease: 70% of the lungs <17 Gy.
- Healthy lungs: 60 % of the lungs <20 Gy.
- Esophagus: Dmax < was 4.0 Gy per fraction.
- Chest wall: <30 Gy to 30 cc and <60 Gy to 3 cc.
- Spinal cord: <2 Gy per fraction and <45 Gy total.



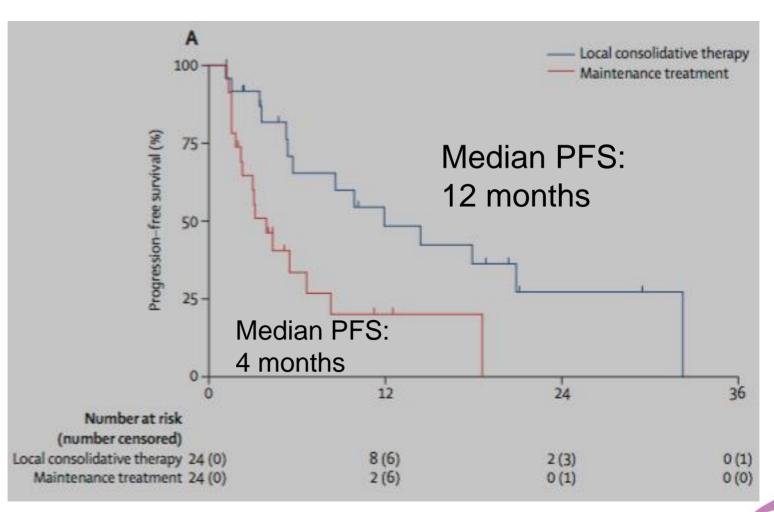


Toxicity (N=28)	Grade I	Grade II	Grade III
Pneumonitis	11	3	1
Chest wall pain	6		
Skin	6		
Esophagitis	3		

Select group of patients that benefit from aggressive local treatment for oligometastatic disease



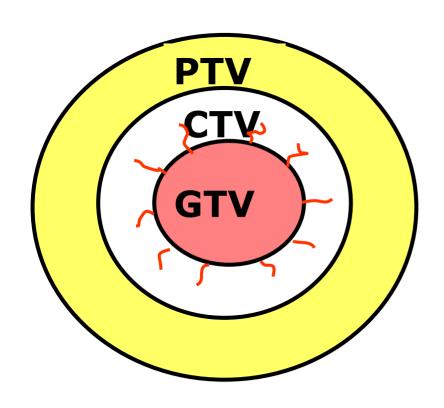
Local consolidative therapy versus maintenance therapy or observation for patients with oligometastatic NSCLC

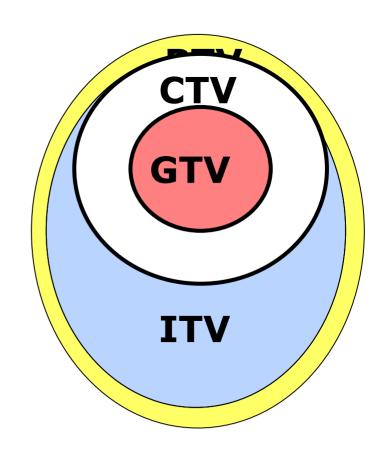


Gomez D et al. Lancet 2016

ICRU 50

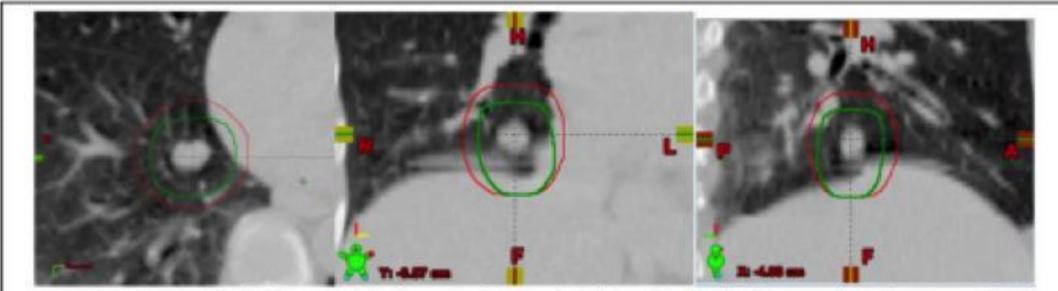
ICRU 62





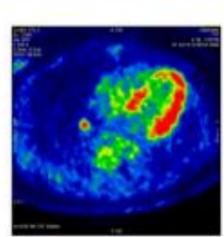
GTV= Gross Tumor Volume, CTV=Clinical Target Volume, PTV=Planning Target Volume, ITV=Internal Target Volume

V. Bettinardi et al./Radiotherapy and Oncology 96 (2010) 311-316

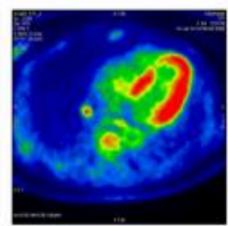


CT Image (Left Transaxial, Center: Coronal, Right: Sagittal).Comparison between PTV_{4D} (green contour) and PTV obtained by standard expansions (red contour)

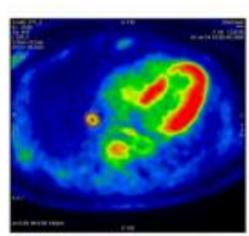




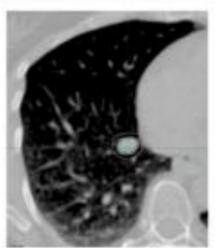
Single 4D-PET Phase



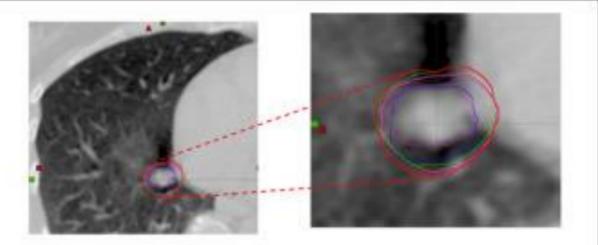
Sum of the 4D-PET Phases



BTV (red contour) also representing ITV_{BTV}

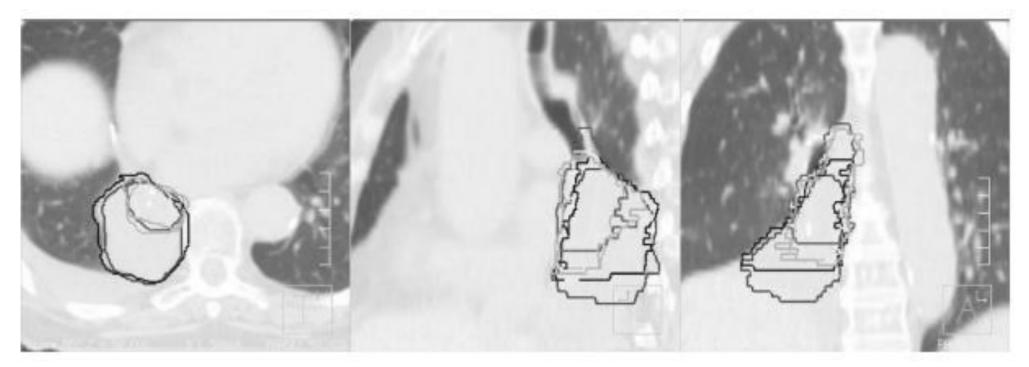


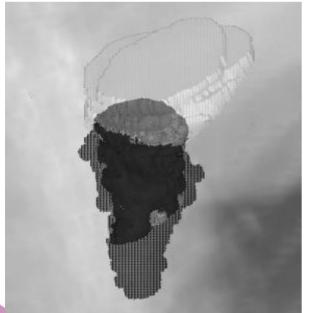
Single 4D-CT Phase GTV (light blue contour) CTV (pink contour)



CTVs from single 4D-CT phases and ITV_{CTV} (red contour) obtained by their convolution (Boolean Union)







Inter-observer and intra-observer reliability for lung cancer target volume delineation in the 4D-CT era





Contents lists available at SciVerse ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



Lung cancer radiotherapy

An evaluation of an automated 4D-CT contour propagation tool to define an internal gross tumour volume for lung cancer radiotherapy

Stewart Gaede a,b,c,d,*, Jason Olsthoorn e, Alexander V. Louie b, David Palma b,c, Edward Yu b,c, Brian Yaremko b,c, Belal Ahmad b,c, Jeff Chen a,b,c,d, Karl Bzdusek g, George Rodrigues b,c,f

Conclusions: Automated 4D-CT propagation tools can significantly decrease the IGTV delineation time without significantly decreasing the inter- and intra-physician variability.



^{*}Physics and Engineering Department; and b Department of Radiation Oncology, London Regional Cancer Program, Canada; Department of Oncology; and Department of Medical Biophysics, University of Western Ontario, Canada; Department of Mathematics, University of Western Ontario, Canada; Philips Radiation Oncology Systems, Flitchburg, WI, USA

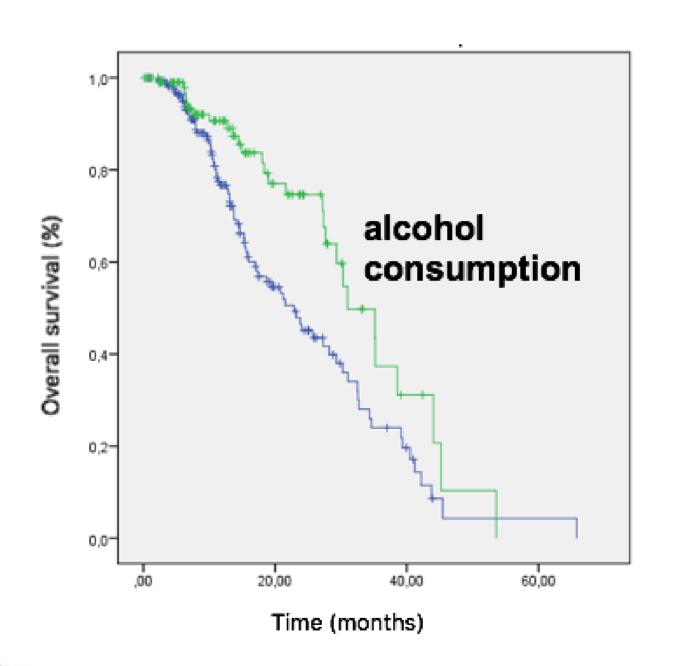
Case 1: Oligorecurrence of lung cancer

- A 65-year-old male presented to the emergency department with a two-week history of upper back pain
- Pertinent social history includes a 34pack year history of tobacco smoking, as well as history of heavy alcohol consumption in the past.
- Chest X-ray and CT scan showed a RUL nodule (14 mm)



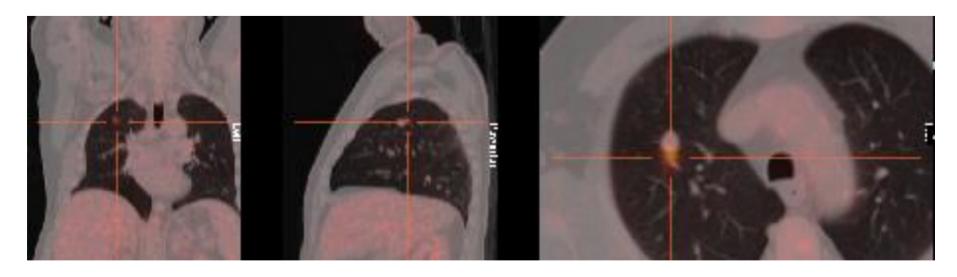


NSCLC in Seville (Spain) 2013-2016





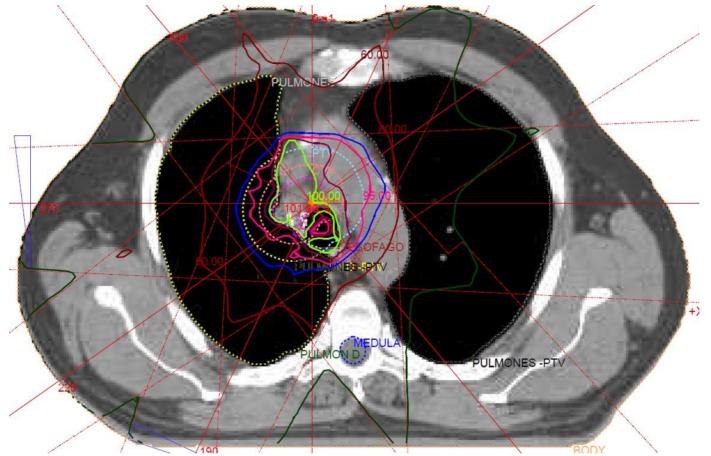
• PET/CT: SUVmax 5,1



- The patient underwent RUL lobectomy and mediastinal lymph node dissection.
- Final pathology report was consistent with high-grade large cell neuroendocrine carcinoma.



• At 2 years follow up, the CT scan showed mediastinal recurrence that was treated with concomitant radiochemotherapy (total radiation dose 66 Gy at 2 Gy/fraction).



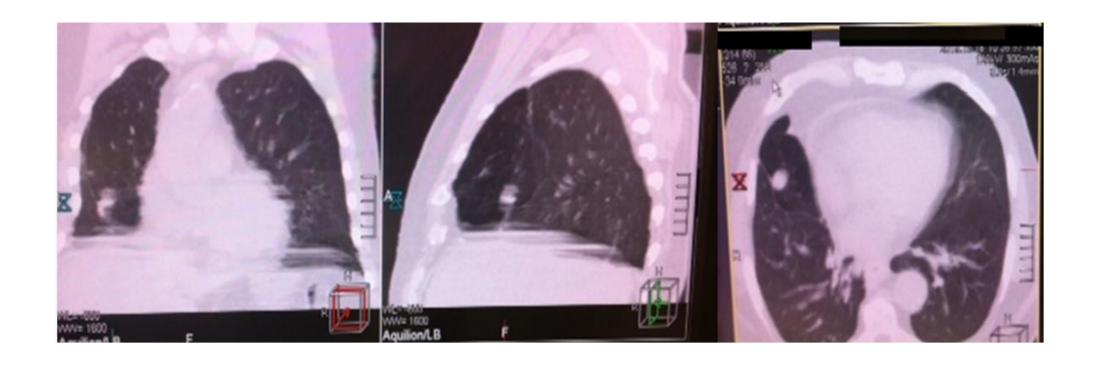


• At 3 years follow up , the CT scan showed a RML recurrence (15 mm nodule).





Motion artifacts are commonly seen with thoracic CT images





Motion artifacts

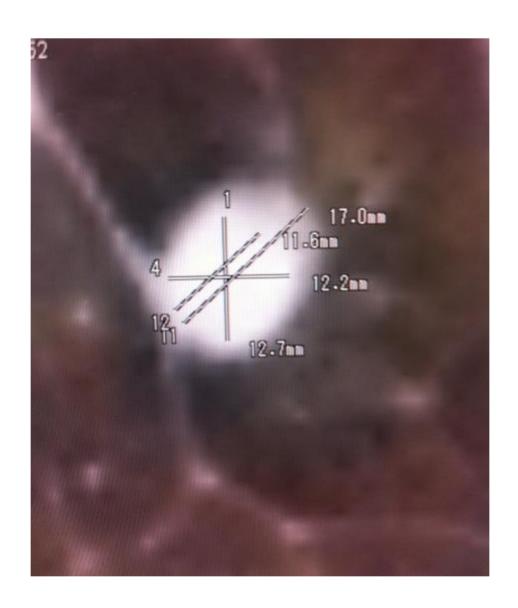








Tumor movement





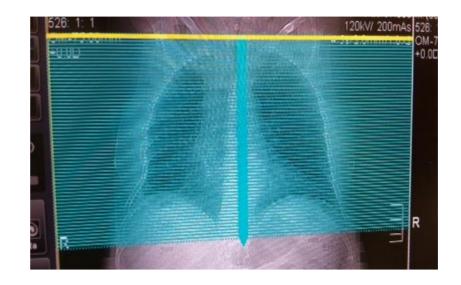
- Diagnosis
- Oligorecurrence of lung cancer
- Treatment
- Radiation Therapy (SBRT)
- Radiation Therapy Dose Prescription:
- PTV (RML nodule): 50 Gy at 12,5 Gy/fraction



Take home message

- Further research is necessary to assess the survival outcome and late toxicity with a longer follow-up for oligometastatic lung cancer
- Different strategies such as 4D repiratory gated acquisition techniques are needed for tumor motion control
- The consecuences of lower doses ("bath dose") in the OAR is still unknown





Questions:

- Immobilization
- Positioning
- Organ at risk contouring
- Set-Up
- Verification
- Radiation technique







Case reports: Lung





Mirjana Josipovic

Dept. of Oncology, Rigshospitalet & Niels Bohr Institute, University of Copenhagen Denmark

Advanced skills in modern radiotherapy June 2017



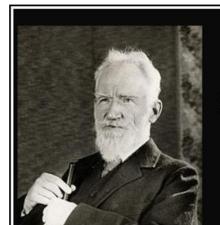
Dose prescription in lung SBRT

Danish national guidelines for lung SBRT

- Prescribed dose: 22 Gy x 3
 - > To the isocenter
 - > PTV encompassed by 15 Gy isodose

Nordic SBRT study group (& Rigs)

- Prescribed dose: 15 Gy x 3
 - To the PTV encompassing isodose
 - ➤ Isocenter dose should be 22 Gy / fraction



England and America are two countries separated by a common language.

(George Bernard Shaw)

Dose prescription in lung SBRT

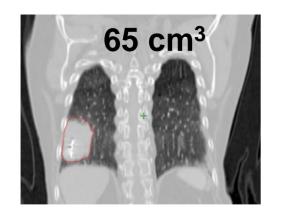
Table 1Patient and treatment details: conformity index (CI) is defined as the volume enclosed by the prescription isodose divided by the PTV volume. Delta⁴ phantom was used to measure gamma agreement index (GAI, 2 mm/3%).

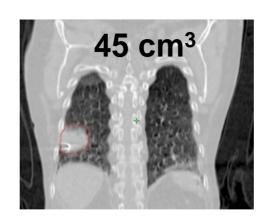
Patients	Tumor site	PTV (ccm)	Treatment schedule	Energy	# arcs	# MU	Mean DR (MU/min)	Max DR (MU/min)	Mean treatment time (min)	Mean overall time (min)	GAI (%)	CI
P1	Lung	15.6	8 × 4.5 Gy @70%	X6FFF	2	1074	851	1200	1.26	18.2 (±4.4)	99.3	1.06
P2	Lung	37.7	10 × 5 Gy @80%	X6FFF	1	1563	1351	1400	1.16	26.8 (±7.1)	97.3	1.21
P3	Lung	14.7	5 × 8.5 Gy @85%	X6FFF	2	2530	1225	1400	2.06	18.1 (±4.8)	98.8	1.19
P4, V1	Lung	37.5	3 × 10 Gy @84%	X6FFF	2	3018	1372	1400	2.20	15.5 (±3.9)	98.7	1.07
P4, V2	Lung	37.7	3 × 10 Gy @84%	X6FFF	2	2309	1346	1.400	1 70	122/121	00.6	1 07
P5	Lung	9.8	4 × 10 Gy @89%	X6FFF	2	3074	138	No	interna	ational		16
P6	Lung	64.0	10 × 6 Gy @85%	X6FFF	3	1129	91	110	mterna	ational	L	14
P7	Lung	18.7	15 × 3.3 Gy @85%	X6FFF	2	902	44			1		23
P8, V1	Lung	14.1	12 × 4 Gy @79%	X6FFF	2	1079	94	con	sensus	on aos	e	47
P8, V2	Lung	5.9	11 × 4 Gy @87%	X6FFF	2	1116	90					09
P9	Liver	36.3	10 × 5 Gy @85%	X6FFF	2	956	59	preso	cription	in SRI	RT	11
P10	Liver	283.5	10 × 4 Gy @80%	X6FFF	2	1114	80	Prest	T P CIO			21
P11	Pancreas	131.8	2 × 6 Gy @78%	X6FFF	1	1258	122					98
P12	Pancreas	24.0	10 × 5 Gy @87%	X6FFF	1	1079	137					31
P13	Lung	27.4	8 × 7.5 Gy @73%	X6FFF	2	1704	83					05
P14	Lung	32.7	12 × 4 Gy @72%	X6FFF	2	1082	5⁴		not y	et		04
P15	Lung	24.4	4 × 12 Gy @71%	X6FFF	3	4017	137		•			00
P16	Liver	42.9	4 × 10 Gy @75%	X6FFF	2	2139	1374	1400	1.55	18.7 (±5.7)	98.4	0.97
P17	Lung	53.8	4 × 10 Gy @85%	X6FFF	2	2263	1355	1400	1.66	17.6 (±3.6)	100.0	1.04
P18	Pancreas	40.9	2 × 6 Gy @76%	X6FFF	2	1760	1056	1400	2.04	18.3 (±4.0)	98.9	1.00
P19	Pancreas	30.2	2 × 6 Gy @75%	X10FFF	2	1851	1842	2400	1.05	17.9 (±1.2)	97.8	0.99
P20	Liver	131.8	4 × 12 Gy @80%	X6FFF	2	3860	1396	1400	2.78	15.3 (±3.3)	99.5	0.98
P21	Lung	19.6	4 × 12 Gy @79%	X6FFF	4	3974	1302	1400	3.01	14.0 (±2.1)	99.4	1.02
P22	Adrenal	53.5	3 × 10 Gy @78%	X10FFF	2	3813	1860	2400	2.03	22.5 (±5.1)	98.9	1.01
P23	Lung	22.6	4 × 12 Gy @84%	X6 FFF	2	5668	1396	1400	3.36	13.5 (±2.7)	97.9	1.20
P24	Adrenal	77.9	4 × 10 Gy @88%	X6 FFF	2	3172	1198	1200	2.65	19.7 (±4.1)	99.6	1.02
P25	Lung	40.7	10 × 5 Gy @79%	X6FFF	2	1364	993	1400	1.38	16.2 (±3.9)	99.2	1.06
P26	Lung	61.7	5 × 7.5 Gy @78%	X10FFF	2	2170	1076	2400	2.02	18.0 (±2.4)	98.3	1.02

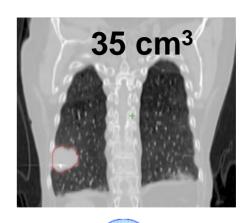


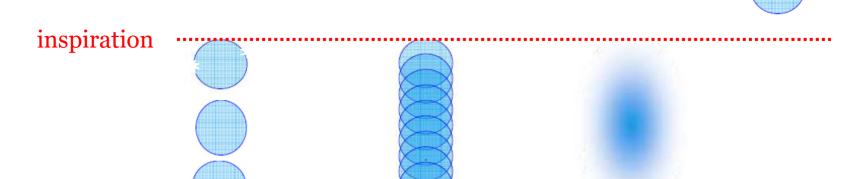
Pre-treatment imaging in lung cancer

What is the true tumour size?









expiration

T

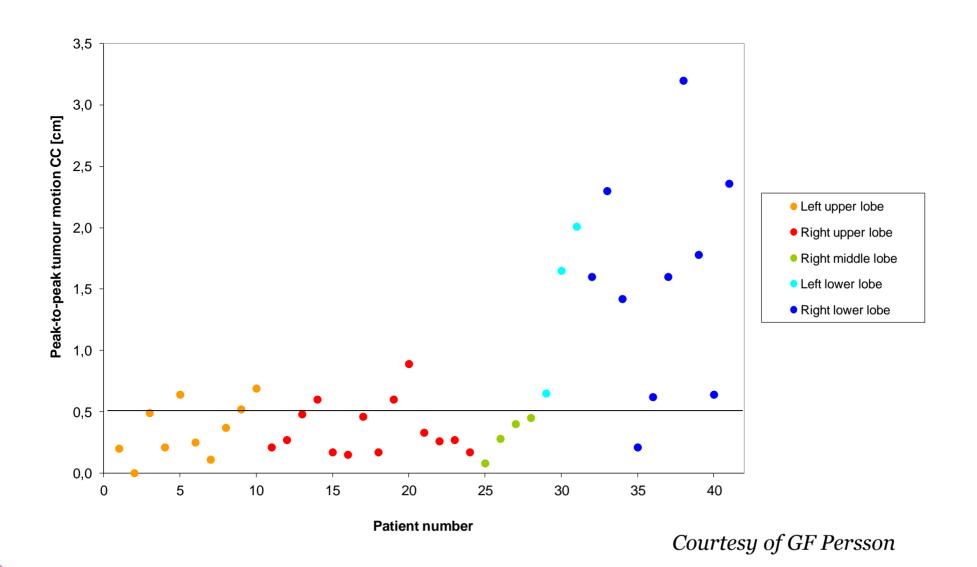
4DCT

PET

Breath Hold

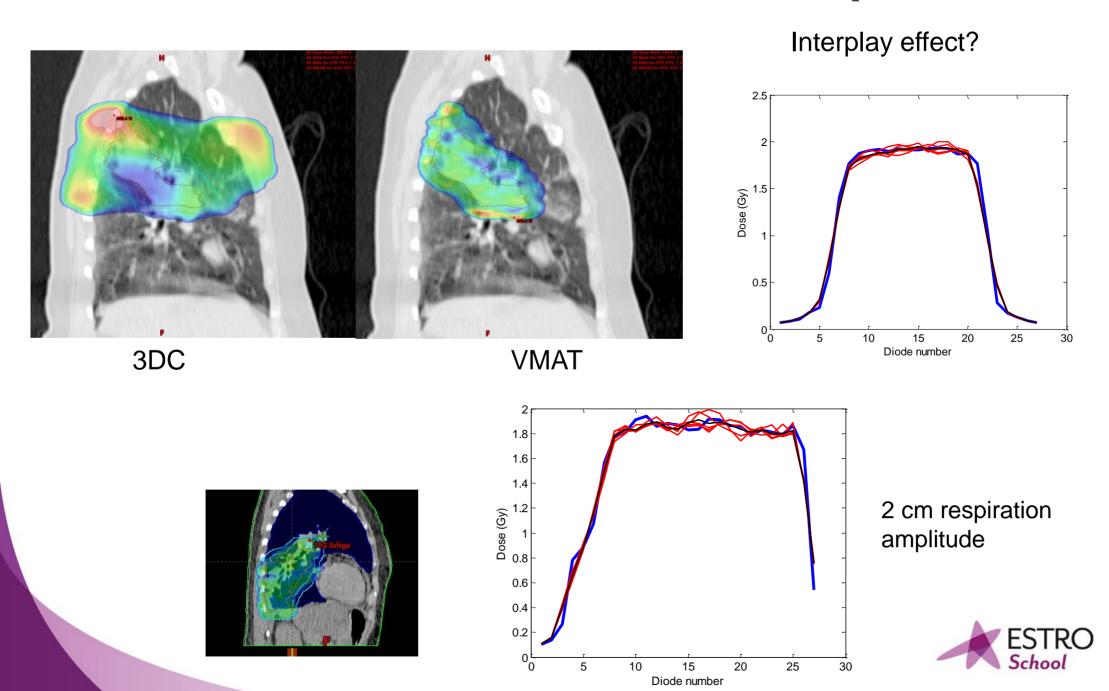


How much do lung tumours move?



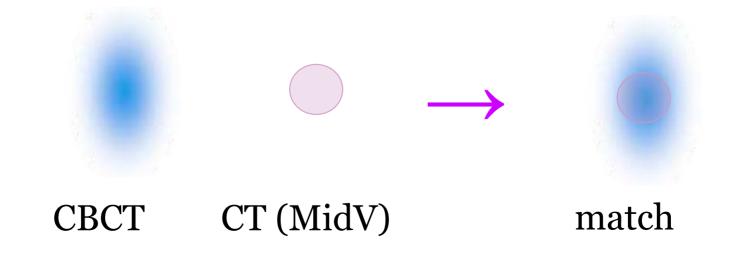


3D conformal vs. VMAT technique



Daily treatment verification

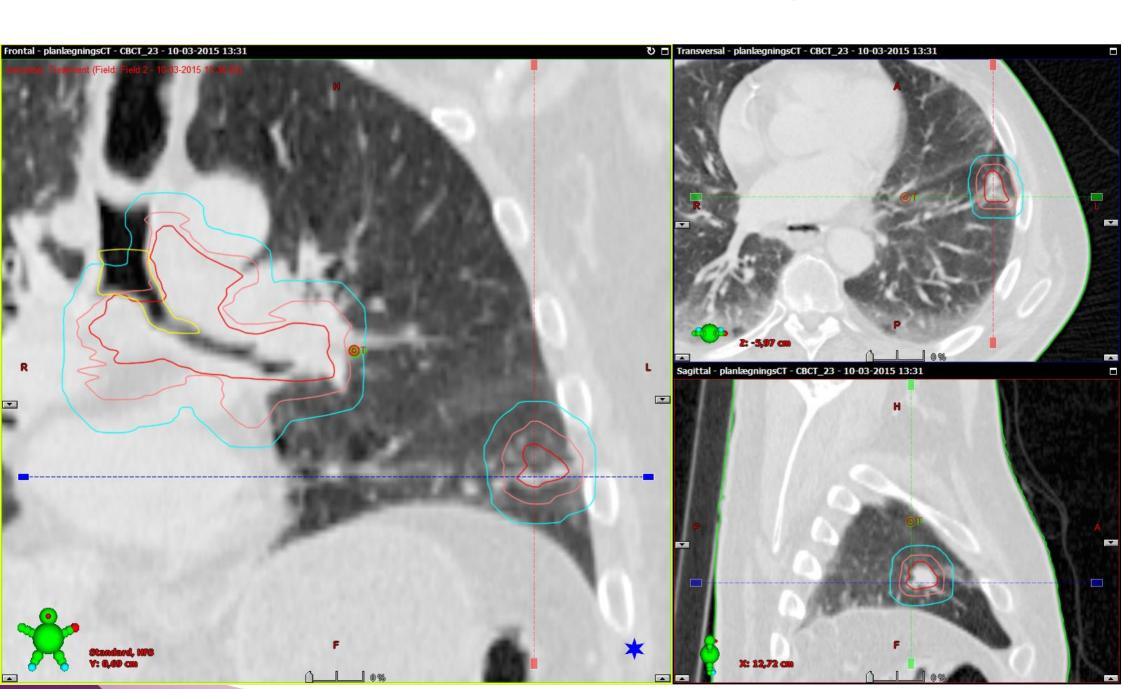
- CBCT acquisition takes ~1 min
 - > Tumour visualisation on CBCT is blurred



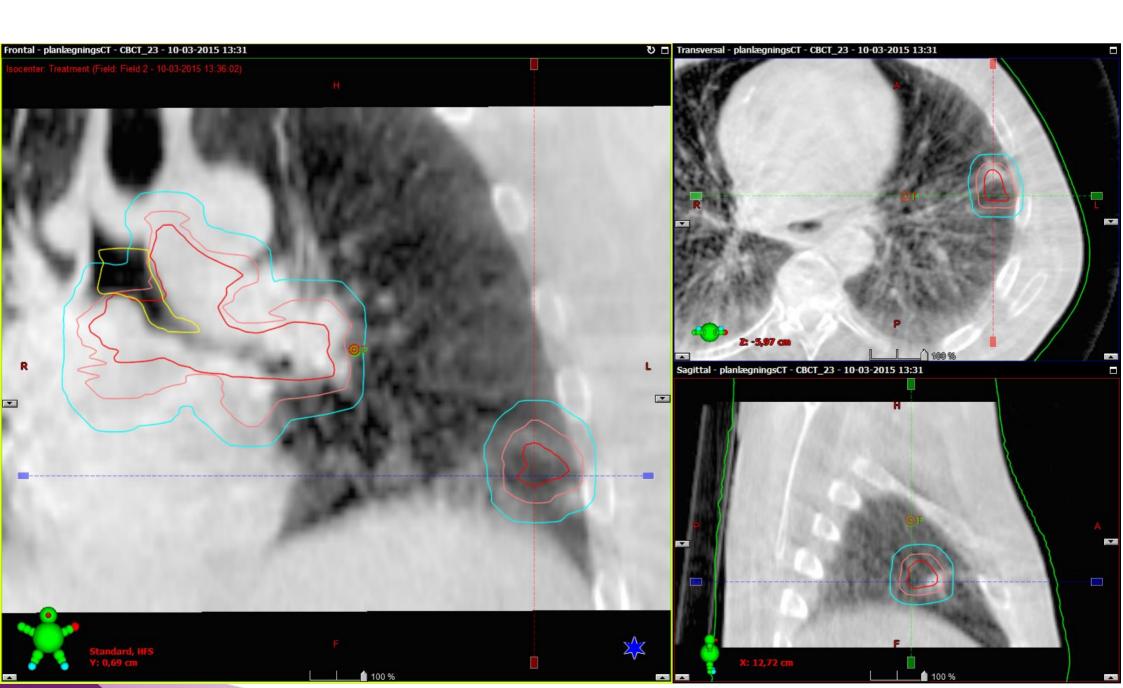
> ...but the image signal (of the tumour) on CBCT is strongest where the tumour is most of the time



Case - locally advanced lung cancer



Case - locally advanced lung cancer



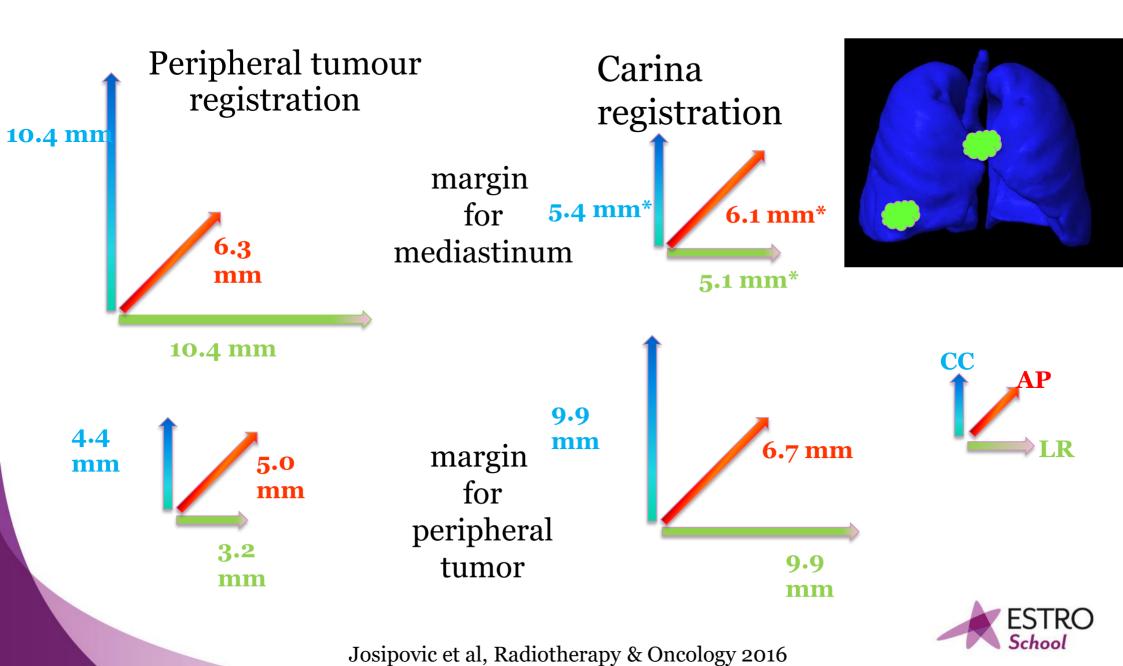
Case - differential motion

The correct answer is...

- What is your IGRT strategy?
- How are your margins designed?
- Is it the first time, (too) large differential motion was observed?



Margins – complex target & daily IGRT



Individual margins

Based on van Herk's margin formula

$$PTV \ margin = \alpha \Sigma + \beta \sqrt{\sigma^2 + \sigma_p^2} - \beta \sigma_p$$

- Respiration amplitude / breath hold reproducibility
- Uncertainties evaluated base on your own data
 - > Tumour baseline shift
 - ➤ 3D instead of 6D match
 - Delineation uncertainty
 - Differential motion
- No. of fractions
- Penumbra width
 - > Lung/bone/softtissue

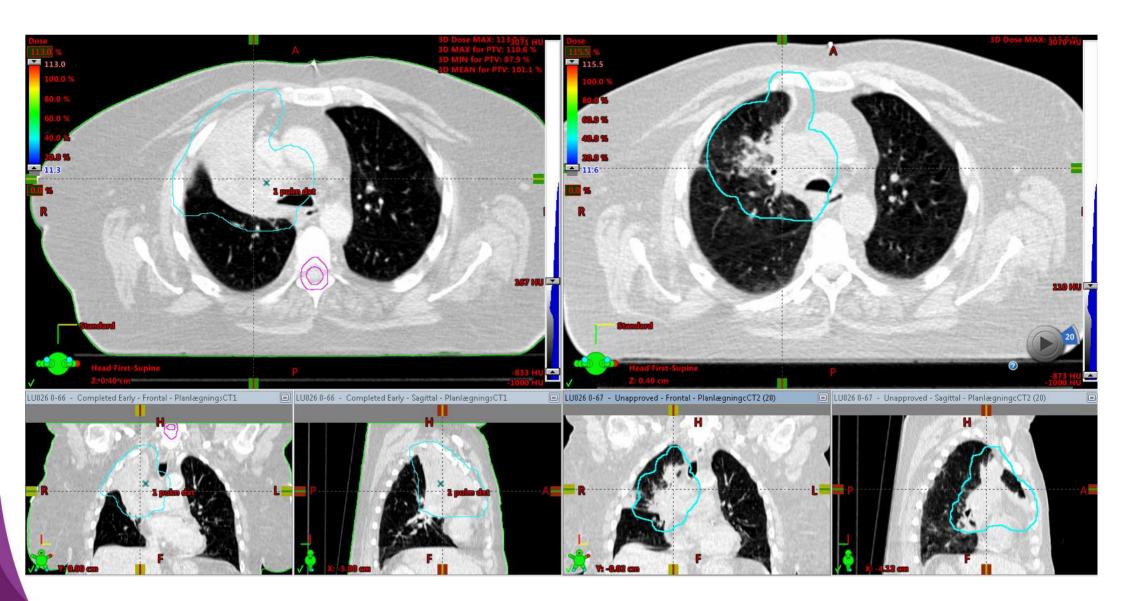


IGRT – uncertainties

- Differential motion
 - Displacement of primary tumour to/from the glands
- Tumour shrinkage during a course of 33 fractions
- Anatomical changes (atelectasis, weight loss, pleural fluid, ...)

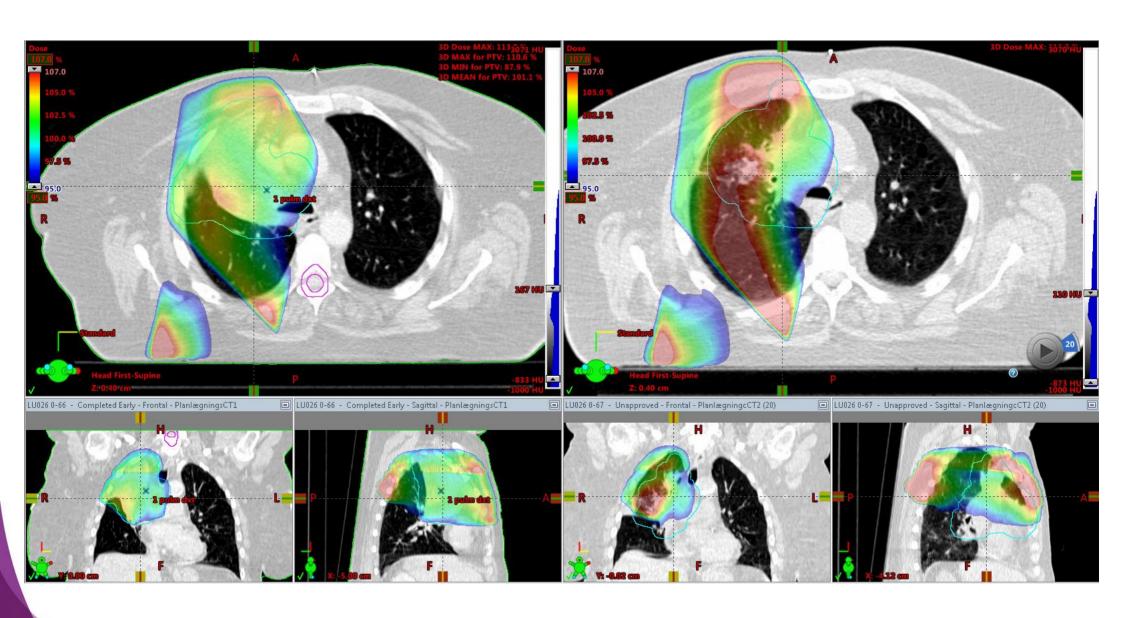


Day 1





Day 1



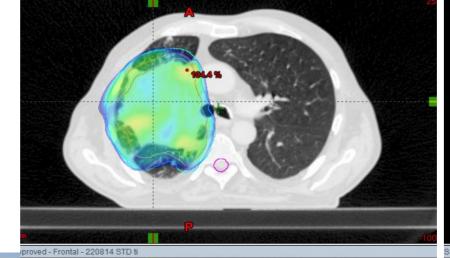
3DC RT

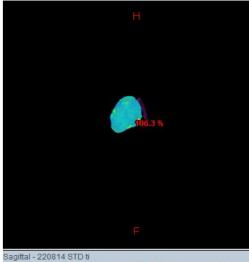
Dmax 107 \rightarrow 115% Dmean 100 \rightarrow 91 %

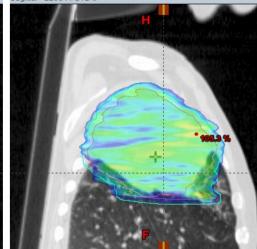


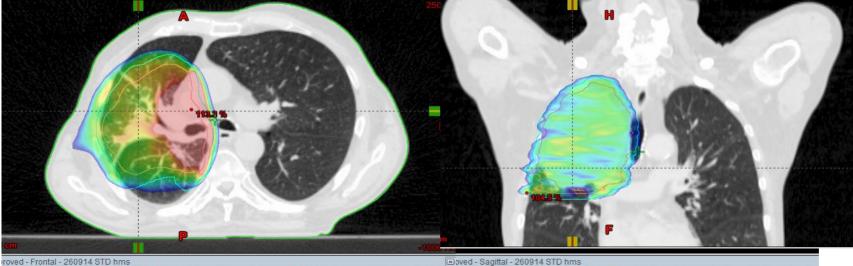
VMAT

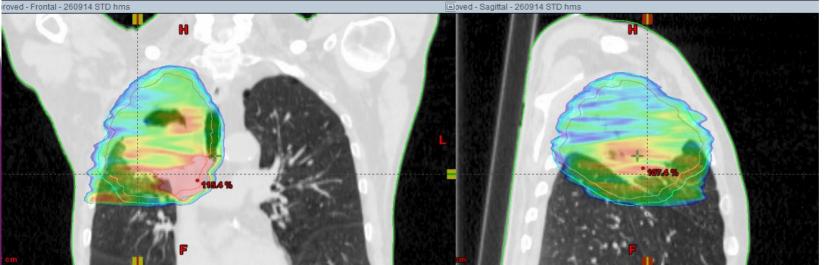
Dmax 106 →117% Dmean 100 → 102.4 %





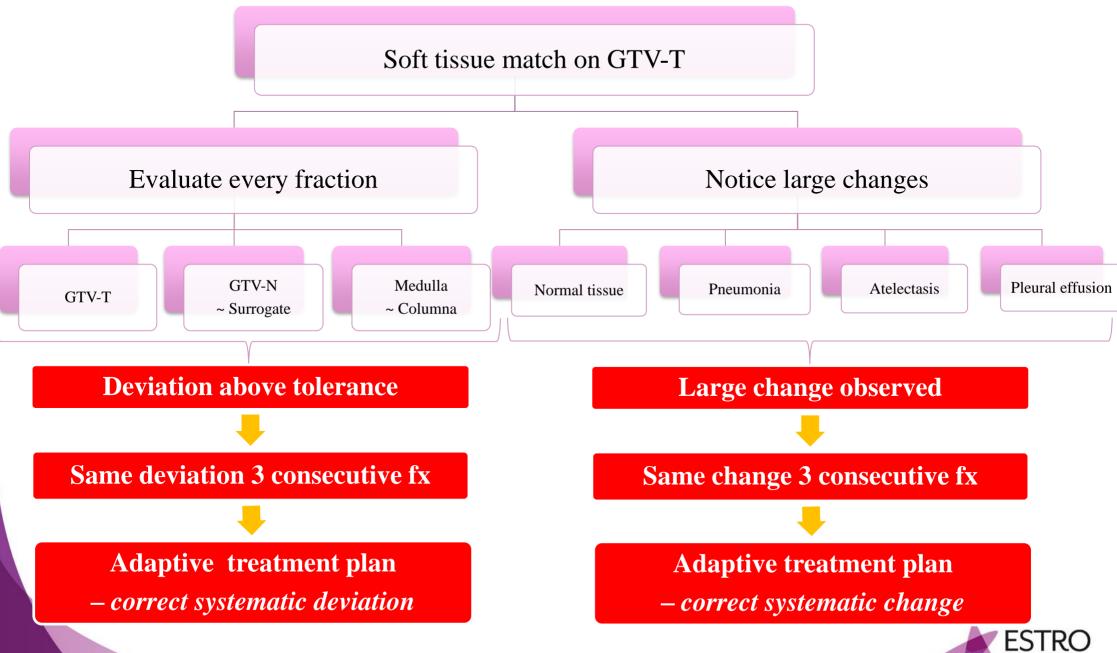








Example of adaptive strategy in lung cancer RT

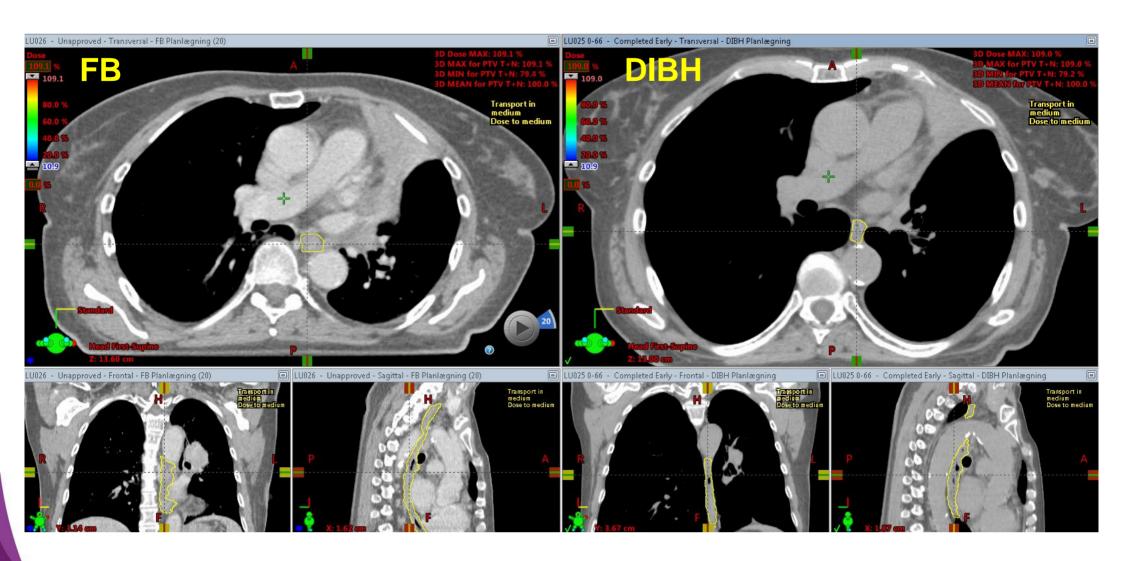


IGRT does not solve all challenges

- When has the anatomical change a significant impact on the treatment
 - i.e. when is it necessary to re-plan the patient?
- How to do the plan adaptation?
 - No published guidelines as yet.
 - Usually, conservative approach with the unchanged CTV delineation is applied



Oesophagus delineation – FB vs DIBH





Oesophagus delineation – FB vs DIBH

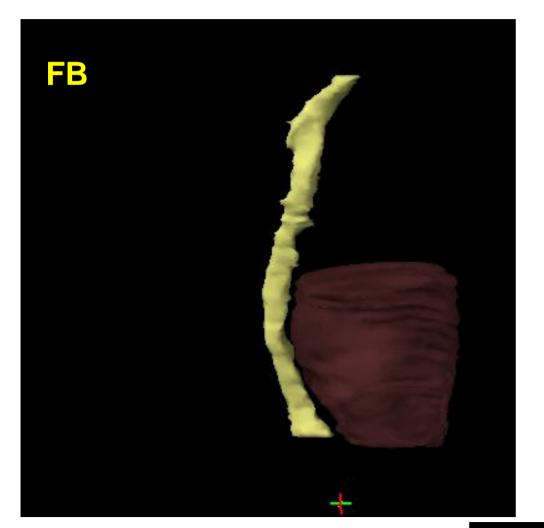




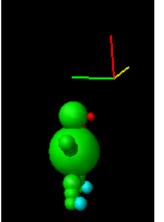
Oesophagus delineation – FB vs DIBH













Lung SBRT on the linac

Martijn Kamphuis MSc Research Radiation Therapist IGRT

Department of Radiotherapy @ AMC Amsterdam, the Netherlands



Key features SBRT

- Reproducable rigid patient fixation
- Managing tumor motion:
 - During imaging
 - During planning
 - During RT
- Very steep dose gradients
- Extreme high BED (100-180 Gy)
 - ➤ 66Gy/2,75Gy→BED 85 Gy
 - > 3*20Gy \rightarrow BED 180 Gy (\triangle 95 Gy)



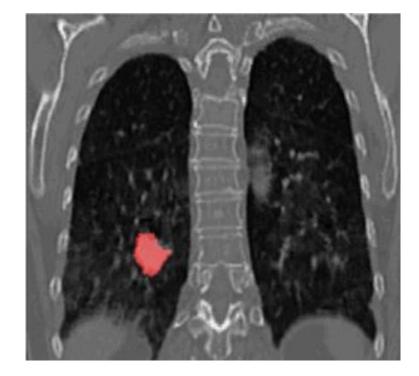
What do we have to manage?

Intra fraction motion

Breathing pattern

Inter fraction motion

Baseline shift



https://www.imi.uni-luebeck.de



Intra fraction motion: amplitude

			Coronal		Sagit	tal
CC	ML	AP	R	L	P	А
4.3 ± 2.4	3.4 ± 1.6	2.8 ± 1.3	1 cm % //	57/3	25%	9 7
(2.6 to 7.1)	(1.3 to 5.3)	(1.2 to 5.1)	1 000	. 1996	200	163
7.2 ± 1.8	4.3 ± 2.4	4.3 ± 2.2	Thom !	1000	3	3
(4.3 to 10.2)	(1.5 to 7.1)	(1.9 to 7.5)	77 73	30/1/10	चर्ची	20
9.5 ± 4.9	6.0 ± 2.8	6.1 ± 3.3	- 2	S S S S S S S S S S S S S S S S S S S	35%	TAN
(4.5 to 16.4)	(2.9 to 9.8)	(2.5 to 9.8)	Mer eng	46	Man A	3. 20
1901 14.40	nor mobility respiration		•	V	1100	

Seppenwolde et.al.



Managing intra fraction motion

Jaches

Active Spanacking

Active Spanacking

Active Spanacking

Active Spanacking

Breathhold

Abdominal compression

ITV concept

Mid-ve Minjana

Mid-ve Minjana

Active Spanacking

Breathhold

Gative

Chasing/tracking

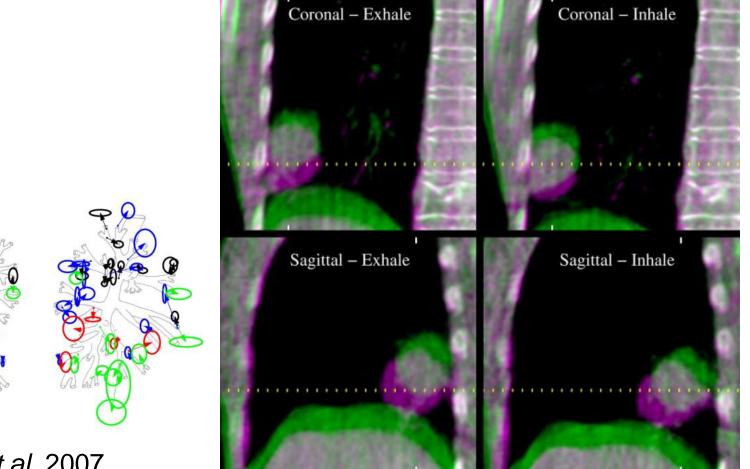


Inter fraction motion: baseline shift





Baseline shift

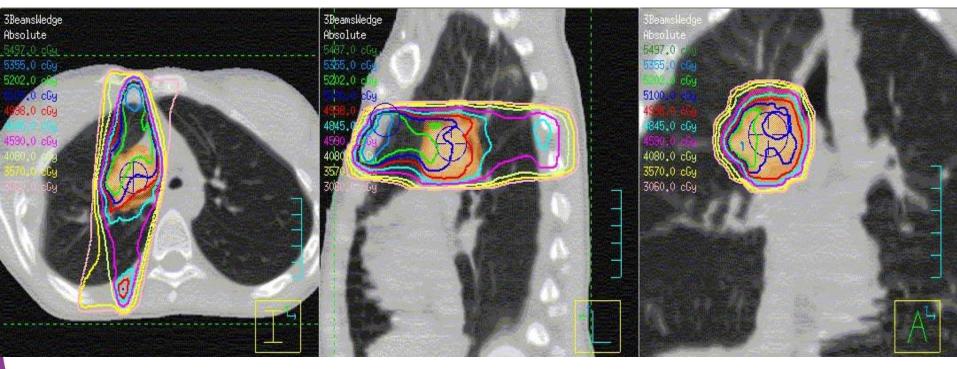


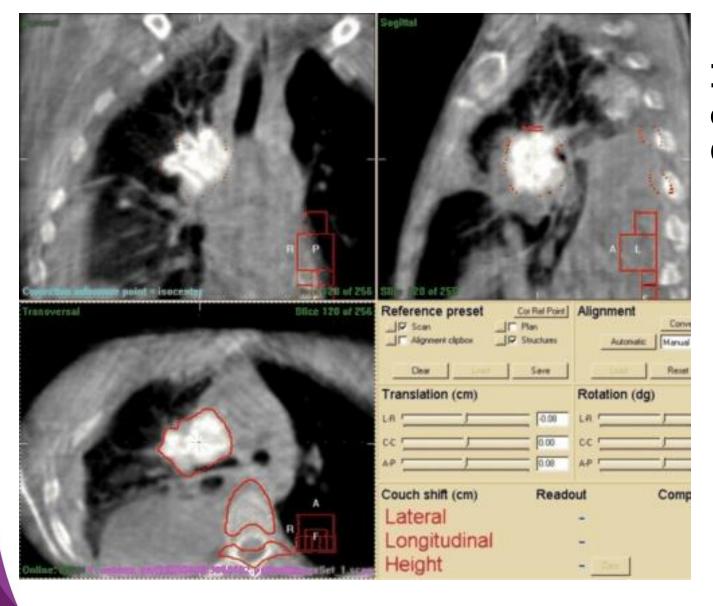
Systematic

J.J. Sonke et al, 2007



Plan: 3 wedged fields, 300 cGy/fx, 5100 cGy total





If corrected online on CBCT



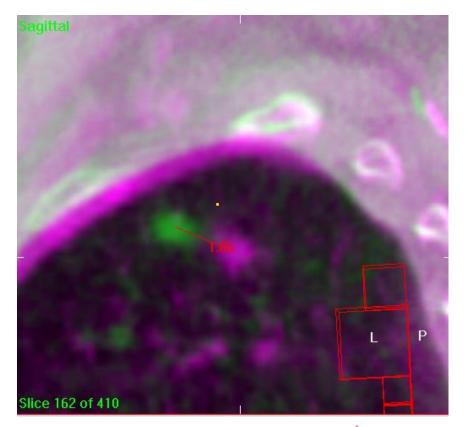
Management of baseline shifts

Introduction of the Planning Risk Volume (PRV)

•Margin around OAR

Combined with:

- 1. Commercial software
- 2. Critical isodose line



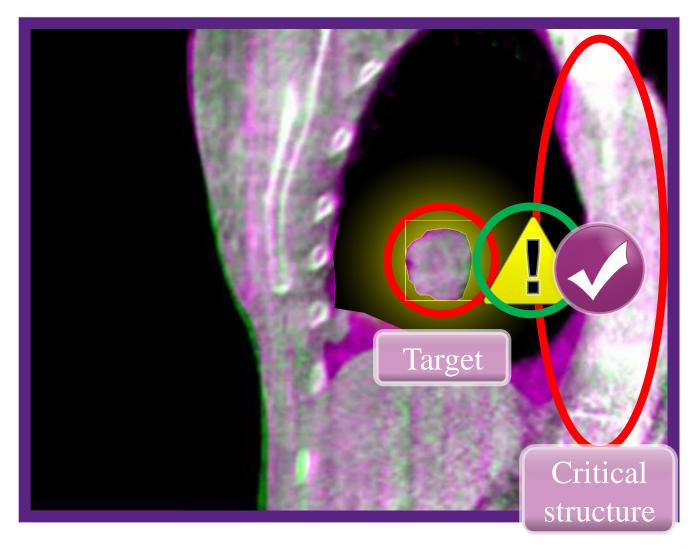


Option 1: NKI/ELEKTA solution

- Image registration on tumor & OAR (dual registration)
- Allowed deviations in distances between tumor and OAR deteremined on Treatment planning
 - Based on dose distributions
- Personal margins are put into the CB-system
 - E.g. 5mm to the Left, 15 mm to the Right



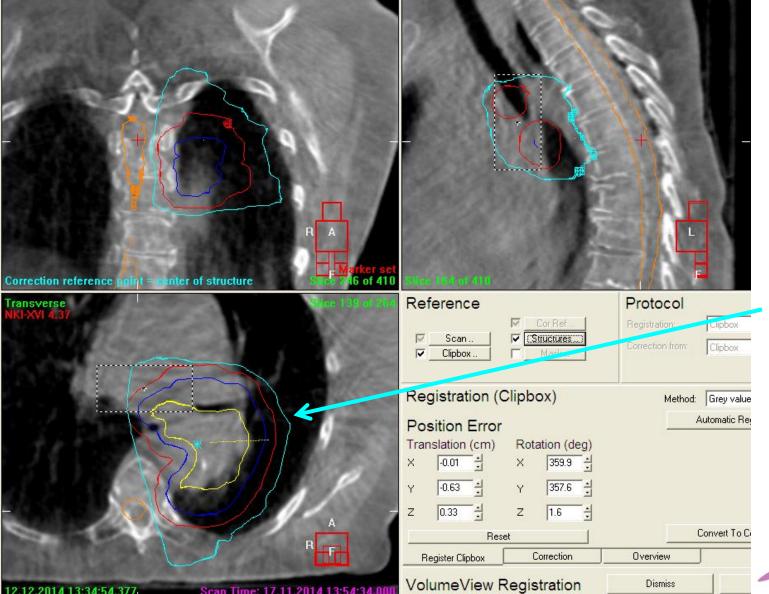
Option 1: NKI/ELEKTA solution



Critical Structure Avoidance (Dual Registration)



Option 2: critical isodose



51 Gy isodose



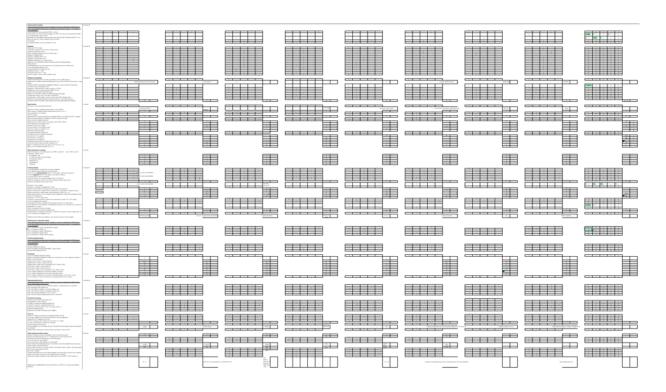


Focus on lung cancer: What a radiotherapy department should offer their patients





ESTRO ACROP guideline development



- Questionnaire of 140 items
- Consensus of 11 experts from the ESTRO SBRT teaching course and their 8 institutions

 ESTRO SBRT

ESTRO ACROP guideline development

Category	Definition:
Mandatory	Minimum equipment and methodology required to achieve clinical outcome in agreement to published prospective clinical trials.
Recommended	Equipment and methodology achieving potentially best clinical outcome and best accuracy currently achievable.
Optional	Equipment and methodology that might improve clinical outcome and accuracy of SBRT without clinical evidence available, yet.
Insufficient	Equipment and methodology resulting in potentially worse clinical outcome compared to published prospective clinical trials.
Discouraged	Equipment and methodology resulting in no improvement in accuracy or clinical outcome and in no other obvious advantage . ESTRO 36 - MATTHIAS

Radiotherapy delivery device

Device	Mandatory	Recommende d	Optional	Insufficient	Discouraged
Conventional C-arm linac	1	0	0	5	2
Conventional C-arm linac with IGRT technology	6	1	0	1	0
Dedicated C-arm stereotactic linac	1	5	1	0	0
Tomotherapy	0	0	6	1	1
Dedicated stereotactic device	0	2	6	0	0

Mandatory:

C-arm linac with CBCT

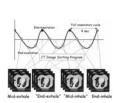
Recommended:

"Stereotactic" C-arm linac - MATTHIAS



Additional technologies

Mandatory



	Mandatory	Recommended	Optional
Respiration correlated 4D-CT	5	3	0

Recommended



	Mandatory	Recommended	Optional
High-resolution MLC < 10mm	2	6	0

Mandatory:

4D-CT

• Recommended:

HR-MLC (5-9mm)

ESTRO 36 - MATTHIAS
GUCKENBERGER
ESTRO

Additional technologies









	Mandatory	Recommended	Optional
Fluoroscopy at simulation for evaluation of tumor motion	0	0	6
Abdominal compression system	0	0	5
Active breathing coordinator system (e.g. ABC system)	0	2	5
Respiration correlated 4D-PET-CT	0	0	8
Implantable fiducial marker system	0	1	6
Implantable transponders e.g. Calypso System	0	0	7
Audio and / or visual breathing motion monitoring system for breathing feedback	0	2	6
Surface Scanner	0	1	5
External breathing motion monitoring system in the treatment room (e.g. RPM system)	0	3	5
Linac with gated beam delivery mode	0	2	6
Flattening filter free (FFF) delivery mode	0	2	6
Very high resolution MLC < 5mm	0	2	6
Robotic 6 degrees of freedom (DOF) treatment couch	1	2	5

Most additional technologies optional



Staffing and Credentialing

		Mandatory
	Written departmental protocol covering all mandatory aspects of SBRT practice	8
	Site-specific SBRT implementation & application based on a multi-disciplinary project team involving Clinicians, Physicists & RTTs	8
FOLLOW UP	Structured follow-up and assessment of clinical outcomes (e.g. local control, toxicity)	8

• Mandatory: Protocols, multi-professional team & structured follow-up



Staffing and Credentialing

	Mandatory	Recommended	Optional
Participation in dedicated SBRT teaching course (e.g. ESTRO SBRT course)	1	7	0
Particpation in Vendor-organized dedicated SBRT training	2	6	0
Supervision of first SBRT treatments by SBRT-experienced colleague	2	5	1
Hands-on training at SBRT-experienced center	3	5	0
External audit of SBRT practice once after implementation	0	4	4
External audits of SBRT practice in regular intervalls after SBRT implementation	0	4	4

Recommended: investment in training and teaching instead of technology

Treatment planning: Planning technique

	Mandatory	Recommended	Optional
3D CRT planning	6	2	0
Dynamic conformal arc planning	2	1	4
Static IMRT planning	0	0	5
Dynamic IMRT planning	0	5	3

Mandatory: 3D-CRT

Recommended: VMAT



Breathing motion compensation

	Mandatory	Recommended	Optional	Insufficient
Population-based margins	1	0	0	4
ITV	7	1	2	0
Midventilation	0	4	4	0
Gating	0	2	6	0
Real-time tracking	0	1	7	0

Mandatory: ITV

Recommended: Mid-ventilation



Image guidance

	Mandatory	Recommended	Optional	Insufficient / discouraged
Stereotactic set-up based on external coordinate system	0	0	2	6
IGRT with Planar EPID imaging only	0	0	0	8
IGRT with Planar kV imaging w/o implanted markers only	1	0	0	7
IGRT with Planar kV imaging with implanted markers only	1	0	6	0
IGRT with Volumetric imaging	6	1	1	0
IGRT with 4D Volumetric imaging	0	7	2	0

• Mandatory:

Recommended:

in-room 3D IGRT in-room 4D IGRT



Thank you for your attention!



'steps' of registration @AMC

Reference settings – 1x

- Visualize your patient in full
- 2. Ask yourself what you are going to treat
- 3. Ask yourself what is the best surrogate for your target volume (within the department protocol)
- 4. Define structures to be displayed, clipbox, corr ref point and algorithm
- 5. Are there patientspecific variables?

Image registration and evaluation

- 1. Visualize your patient in full in green/purple overview
- 2. Automatisch registration (match)
- 3. Evaluation of your registration (match)
- 4. Evaluation of CTV coverage target within PTV?
- 5. Are the rotations within tolerance
- 6. Are there any changes in patient anatomy that might influence the dose distribution
- 7. Evaluation the correction without rotations target within PTV?

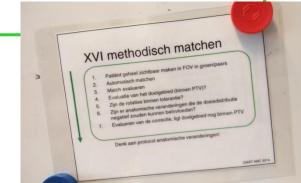


Image registration and evaluation

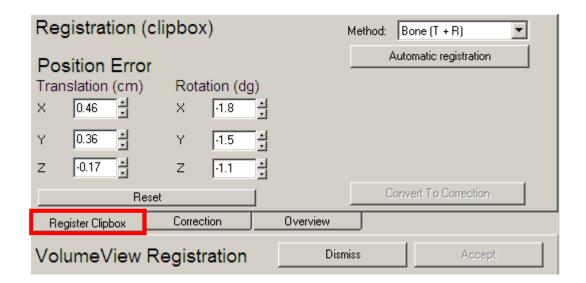
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- Evaluation of your registration (match)
- 4. Evaluation of CTV coverage target within PTV?
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Image registration and evaluation

- 1. Visualize your patient in full in green/purple overview
- 2. Automatisch registration (match)
- 3. Evaluation of your registration (match)
- 4. Evaluation of CTV coverage target
- 5. Are the rotations within tolerance
- Are there any changes in patient and the dose distribution
- 7. Evaluation the correction without rota

Fr.	Datum	PT	V?	Kut	atie	(cm)	(cm)
		JA	NEE	<4°	≥4°		
1							
2							
3							
4							
5							

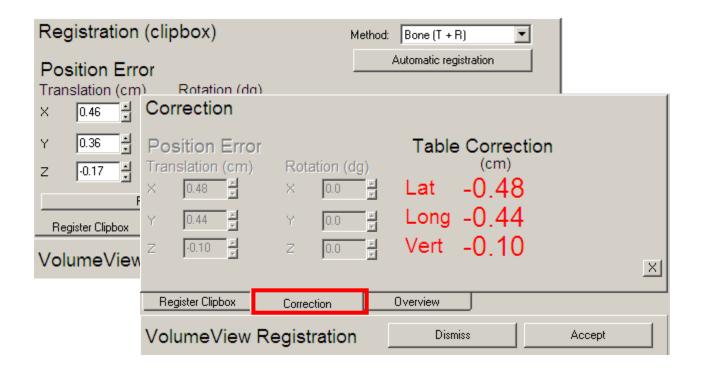
Symantics @Elekta



Tab Register Clipbox:

Full registration in translations and rotations.

On this tab you check the registration and evaluate patient anatomy

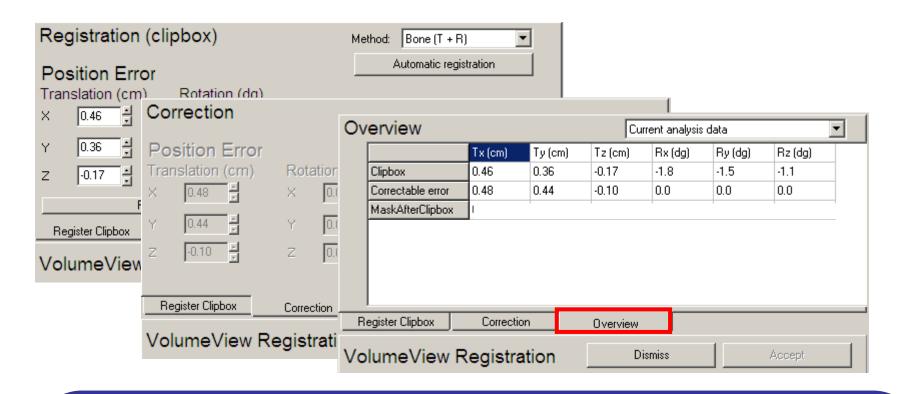


Tab Correction:

Displays the correctable error. It is the registration where the rotations have been recalulated. The registration ourcome is grey because you cannot change the registration in this tab.

In red it displays the table shift to get to the correctable error. In the viewer the position you are going to treat the patient.

Notice the opposite direction of the numbers between correctable error and table shift in x and y.



Tab Overview:

Display an overview:

Full match in translations en rotations: Clipbox

Correctable error: Correctable error

This tab does not display the table shift!!

You can only find that in the tab Correction.

Algorithms @Elekta

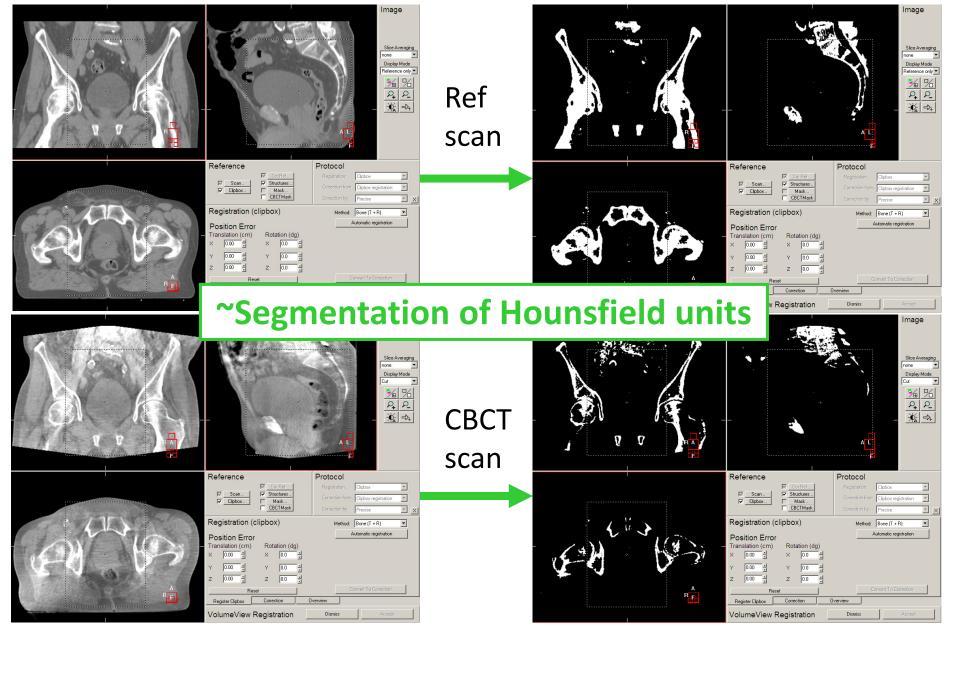
Algorithms

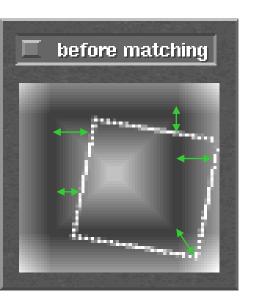
Bone & Seed: chamfer match

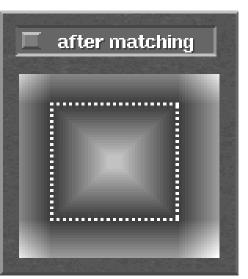
T+R = Translations and rotations

T = Translations only

```
Manual
Bone (T + R)
Bone (T)
Seed (T + R)
Grey value (T + R)
Grey value (T)
Grey value 4D (T)
```





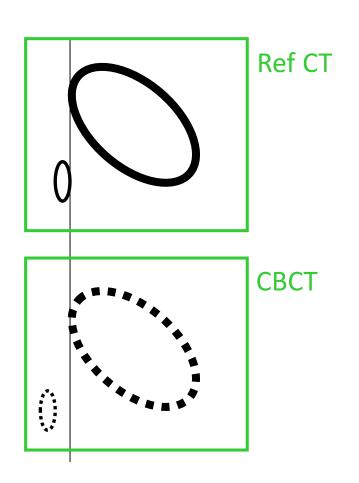


Bone and seed match:

Registration quality is measured by the average distance between the two segmented objects.

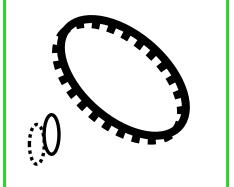
Is quick and reliable. The algorithm indicated whether the outcome was perfect or not 'match may be inaccurate'

Bone and seed algoritme



Pitfall:

If the object is not rigid and not equally balanced in size, the biggest part has more importance.



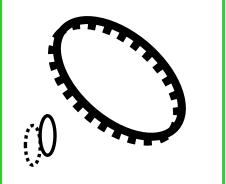
Chamfer match

Bone and seed algoritme

Pitfall:

If the object is not rigid and not equally balanced in size, the biggest part has more importance.

Always visually check whether you got your desired answer

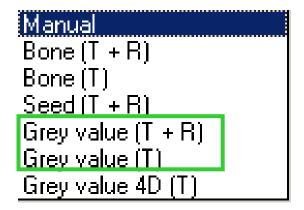


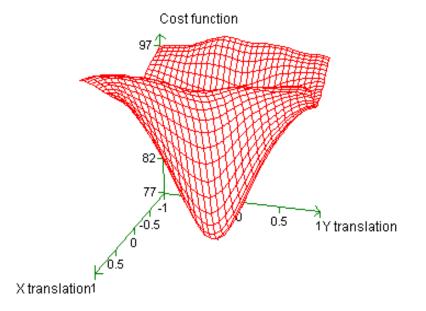
Chamfer match

Algoritmes

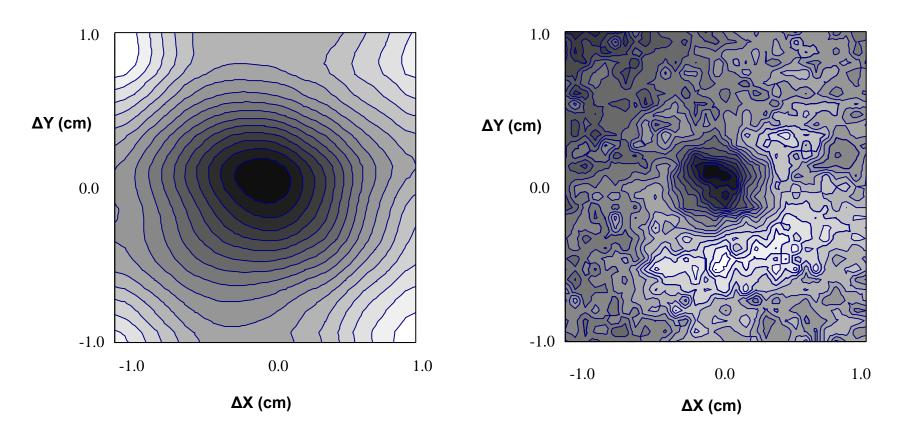
Grey value:

Takes all the pixel values into account:





Cons: Local minima: specifically a problem for small regions of interest depending on cost function



Pitfalls:

If the object is not rigid and not equally balanced in size, the biggest part has more importance.

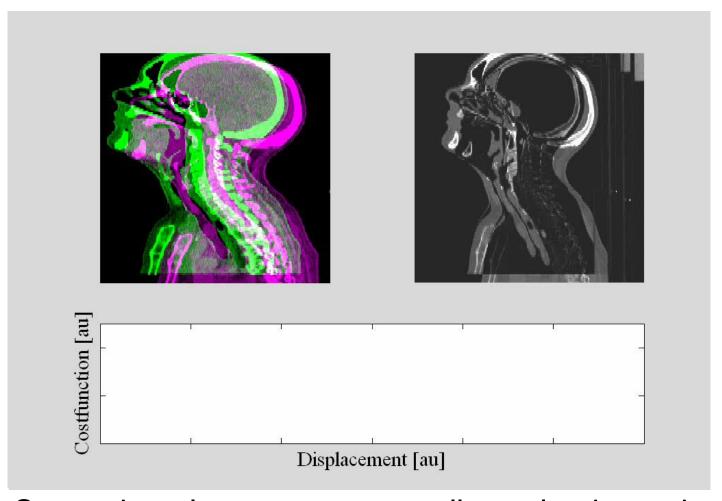
High densities have more influence

Big densities drops have big influence

Artefacts!!

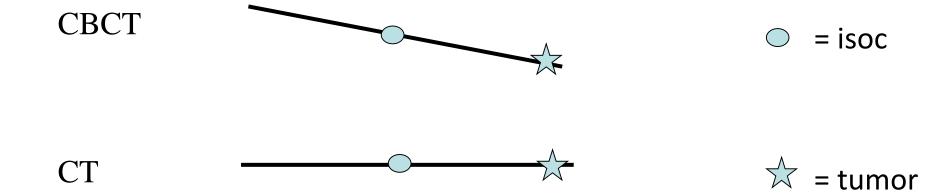
Always visually check the outcome!

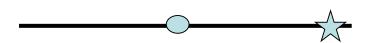
Uses all pixel values in ROI: e.g., sum of squared differences



Somewhat slower to process all voxels: depends on the size of the ROI

CRP @Elekta

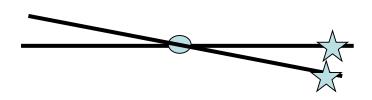




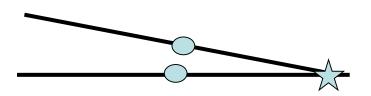
CBCT registered (matched) to reference CT:

Perfect! With 6 degrees of freedom (dof):

3 translations and 3 rotations



After 'convert to correction' with correction reference point at isocentre: isoc is perfecty registerd, tumor is not.



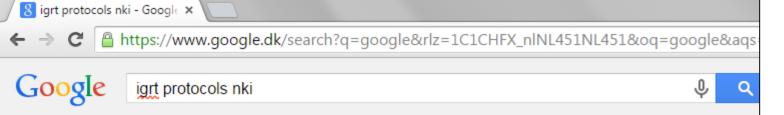
After 'convert to correction' with correction reference point in the tumor: tumor is perfectly registered, isoc is not.

For lazy people only (C)



You can already think IGRT at Treatment Planning! Choose your isocentre at your rotation point of preference (= centre of gravity of target volume (PTV)) Be smart! Nork for you!!!

Let the system work for you!!



Web Afbeeldingen Video's Nieuws Meer ▼ Zoekhulpn

Ongeveer 1.930 resultaten (0,70 seconden)

Tip: Alleen in het Nederlands zoeken. U kunt uw zoektaal instellen in de Vo

[PDF] XVI Engelse Protocols 16_7_2014 - Antoni van Leel www.avl.nl/.../xvi_engelse_protocols_16_7_2014.pd... ▼ Vertaal deze In this document you will find the current IGRT protocols that are used at the questions, you can address them to our imaging RTT's at imagingrt@nki.nl.



XVI Protocols: Netherlands Cancer Institute The Netherlands

July 2014



Contact: Imaging RTT imagingrt@nki.nl

Break-up sessions

Image Registration and Evaluation

Demo Database XVI - Elekta

ESTRO c duras Advance d Skills for Treatment Delivery Ameterds in 2014

> Images Pater Berneller, NO / 8d. Stangage de Jong AVIC

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MATCH PARAMETERS.

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Correction reference point

- Repression -Borganstony -B-D-Dears So or trien assumption
- 4 Correction - Only management - Pirecticly large explina.

Match procedure

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- Receive server of m/a anya point.

r. Gerection - Dry translations







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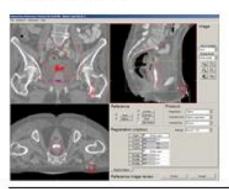
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Emma Endametrium

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Image Registration and Evaluation: Part 2 CBCT (Varian)

Liz Forde, RTT
Assistant Professor
The Discipline of Radiation Therapy
School of Medicine
Trinity College Dublin



Learning Outcomes

- Identify the key features of the Varian OBI system
 - > 2D and 3D image acquisition, registration and verification
- Outline the CBCT acquisition, registration and evaluation process
- Discuss what influences CBCT image quality
- Identify appropriate match structures for the main tumour sites
 - ➤ kV 2D/2D and CBCT
- Discuss possible clinical scenarios that require troubleshooting

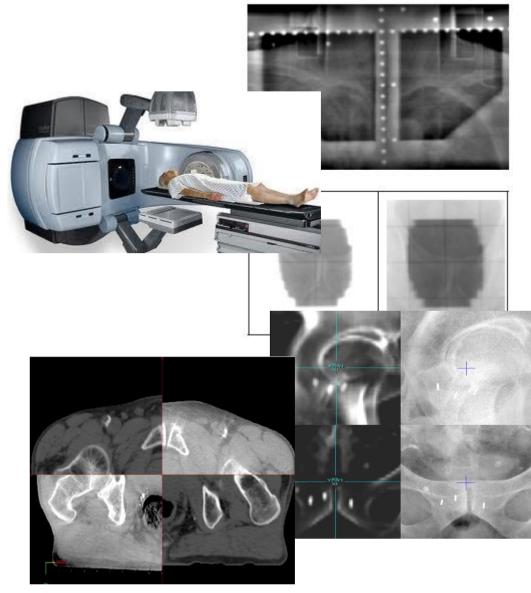






Key Features of Varian OBI

- 2D
 - > MV and kV
- 2D/2D
 - > MV and kV
- 3D
 - > kV
- Fluoroscopy (2D + time)
- Remote couch shift









Key Features of Varian OBI

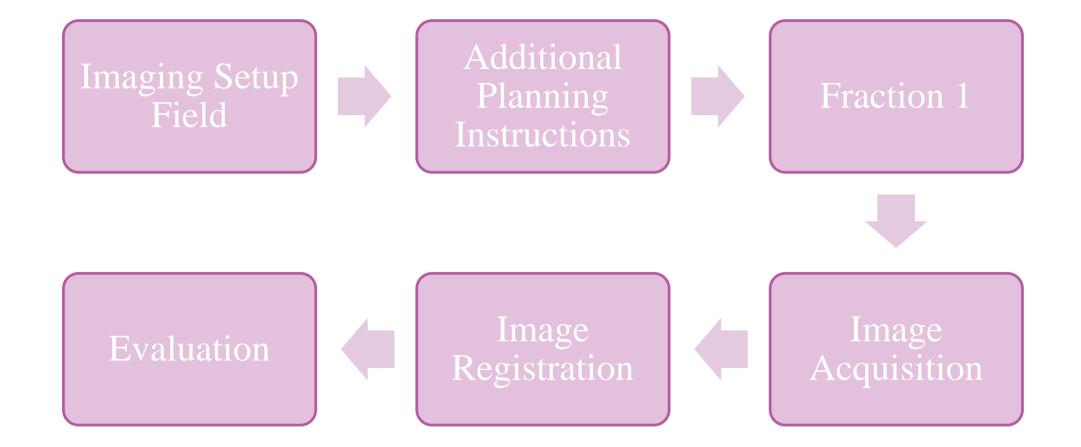
	Standard- Dose Head	Low-Dose Head	High- Quality Head	Pelvis	Pelvis Spotlight	Low-dose Thorax
X-Ray Voltage [kVp]	100	100	100	125	125	110
X-Ray Current [mA]	20	10	80	80	80	20
X-Ray Millisecond [ms]	20	20	25	13	25	20
Gantry Rotation Range [degrees]	200	200	200	360	200	360
Number of Projections	360	360	360	655	360	655
Exposure (mAs)	145	72	720	680	720	262
CTDIw (mGy / 100 mAs)	2.7	2.7	2.7	2.6	2.0	1.8
Dose (cGy)	0.39	0.2	1.94	1.77	1.44	0.47
Fan Type	Full Fan	Full Fan	Full Fan	Half Fan	Full Fan	Half Fan
Bow Tie Filter	FULL	FULL	FULL	HALF	HALF	HALF
Default Pixel Matrix	384 x 384	384 x 384	384 x 384	384 x 384	384 x 384	384 x 384
Slice Thickness [mm]	2.5	2.5	2.5	2.5	2.5	2.5
Reconstruction Filter	Sharp	Standard	Sharp	Standard	Smooth	Standard





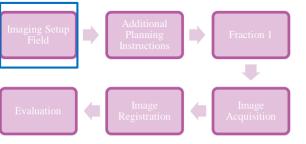


The IGRT Process

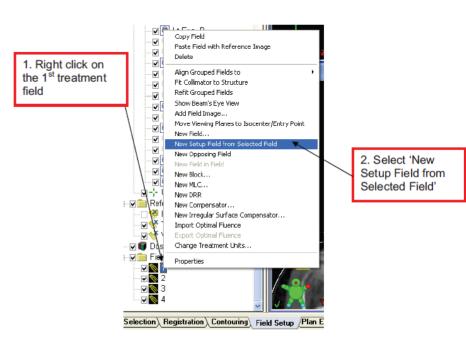


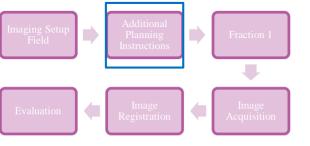






- Create setup fields in planning
- Consider the position of the isocentre
 - Varian does not have a "Correction reference point"
 - > IMRT and VMAT are forgiving with isocentre placement
 - CBCT may need to shift laterally for clearance
 - You will be prompted on the linac





Treatment Area

All treatment areas

Imaging Type

PTV

CBCT, kV or MV

IGRT Setup in Planning

• Additional contours to be outlined and/or "sent across" for image verification

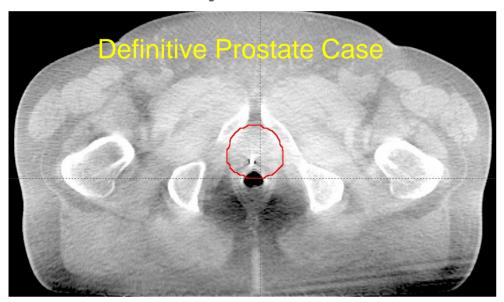
Extra Contouring

All treatment areas	CBCT	FSD tolerance rings				
		(see site specific planning protocols fo	r			
		instructions and size of the rings)				
Chest	kV or MV or CBCT	Carina				
Abdomen	kV or MV or CBCT	Carina				
Breast/Chest Wall	MV	Lung (treatment side) and Body				
Prostate	CBCT	Convert dose to structure				
		(see prostate protocol for instructions)				
Post-Prostatectomy	CBCT	Convert dose to structure	Approval			
·		(see prostate protocol for instructions)	approval 2d Structures in Reference Images	Actual SSD		
		Select the structures to be projected	Rt Eye_P IMRT PTV 63_P Lt Eye_P Pit-Optic Chiasm Spinal Cord_P Lt Lung Body Rt Lung Select All	Field ID Planned Actual 1 94.5 94.5 9 2 93.5 93.5 9 3 95.0 95.0 9 4 95.3 95.3 9 52 94.5 94.5 9 53 94.4 94.4 9 51 94.4 9 58 94.4 9 68 96 96 96 96 96 96 96 96 96 96 96 96 96		
		2. Select 'Next'	- DRRs Generate DRRs to Fields -Field Splitting ■ Split large IMRT fields in Eclipse	Treatment time Calculate Treatment Times Multiply with Factor		

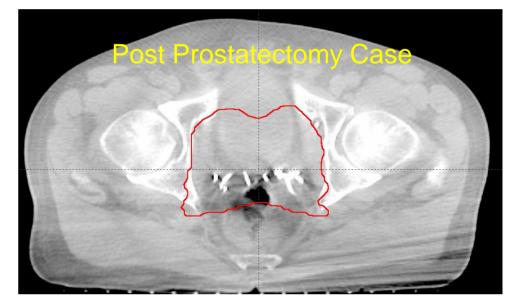
 \underline{N} ext \geq

- Additional contours to be outlined and/or "sent across" for image verification
 - > In Field Setup (Eclipse TPS) "Convert isodose line to structure"

80Gy isodose line



68Gy isodose line

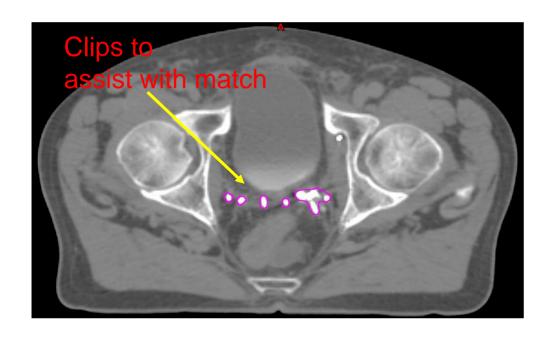








- Additional contours to be outlined and/or "sent across" for image verification
 - ➤ In Contouring Workspace in Eclipse TPS



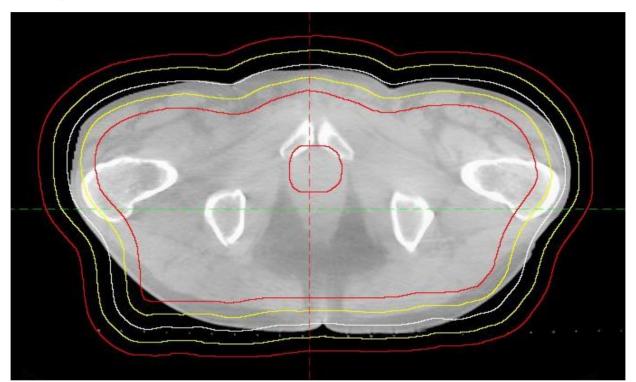








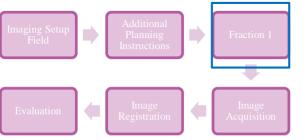
- Additional contours to be outlined and/or "sent across" for image verification
 - In Contouring Workspace in Eclipse TPS "Wall Extraction" tool from Body contour











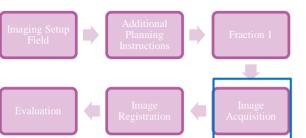
Fraction 1 Considerations

- Clearance
- Education
 - ➤ Who should be present for first day scan?
 - > RO, MP, RTT responsible for plan, Senior RTT
- Documentation!
 - Anything weird and wonderful
 - Structures to include/avoid
- Set VOI box and decide on additional registration variables
 - > This will ensure consistency throughout the course







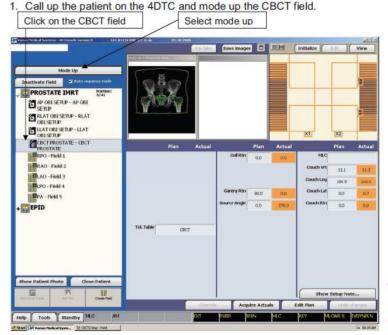


The Image Acquisition Process - CBCT

- Select correct bow tie filter for treatment site
- 2. On fraction 1 consider checking rotation/clearance whilst in room
- 3. Mode up CBCT setup imaging field
 - 1. Note this is incorporated in the individual patient's plan

Scan Name	Gantry Rotation Required	Bow-tie Filter Required	Treatment sites to be used on	Field of View
Standard- dose head	200	Full	-	24cm
Low-dose head	200	Full	-	24cm
High-quality head	200	Full	Head & Neck Brain	24cm
Pelvis	360	Half	Pelvis (includes: Prostate Rectum Bladder Gynecological)	45cm
Pelvis spotlight	200	Full	-	24cm
Low-dose thorax	360	Half	Chest Abdomen	45cm

NB: See pictures below to distinguish between the bow-tie filters





The Image Acquisition Process - CBCT

- 4. Select 3D/3D match
- 5. Acquire new scan
- 6. Complete details
 - 1. Slice thickness
 - 2. Orientation
 - 3. Full fan or half fan
- 7. Start scan
- 8. Accept and export











CBCT Image Quality

- What impacts on image quality?
 - CBCTs use a large flat panel detector increases scatter
 - Permanent anti scatter filter built into detector panel

Scatter decreases image contrast, increases noise, possible registration errors and also patient dose

CT Numbers (HU) affected







CBCT Image Quality

Machine characteristics

- > MV or kV
- > Acquisition time
- > Scan length
- Filters used
 - Bow Tie filter added to source panel







Bow Tie Filters

- Decrease patient dose
- Two types used in different modes: Full fan or half fan mode
- Full fan mode: image is acquired at the central axis on the detector panel and images acquired from 200° rotation
- Half fan mode: the detector is offset laterally acquiring only half of the projection of the patient
 - Detector panel is offset laterally, rotates a full 360° captures only half a projection and reconstructs the image from that
 - Recommended for larger FOV (pelvis)
 - Half fan filters result in the greatest HU discrepancy b/w CT and CBCT (Ding *et al.*, Yoo and Fang-Fang, Seet *et al.*)







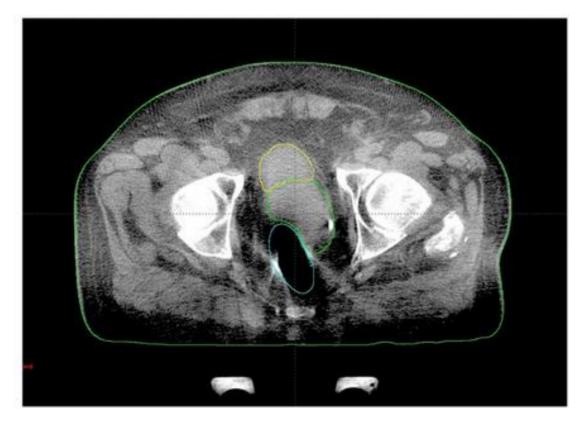
CBCT Image Quality

Patient characteristics

- Size
 - Poor image quality as the patient contour approached the limits of the FOV
- > Tissue heterogeneity
- High dense structures
 - Hip prosthesis
- Motion
 - Increased risk of motion with slow scan time
 - E.g. peristalsis, breathing and gas



CBCT Image Quality



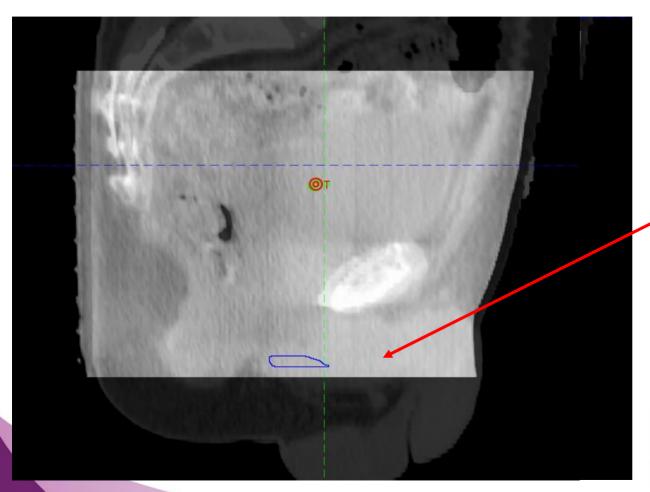
Reggiori et al., 2010

Degradation of image quality due to patient size and gas passing through rectum at time of scan



The Image Acquisition Process

- Make sure you image what you need to match and review to
- Option to offset the couch to ensure appropriate anatomy is visualized



Definitive Prostate Case

Couch now offset to include Penile Bulb in image







The Image Acquisition Process

 Option to offset the couch to ensure appropriate anatomy is visualized

Missing Superior PTV



Contents lists available at ScienceDirect

Radiotherapy and Oncology

journal homepage; www.thegreenjournal.com



Review

The European Society of Therapeutic Radiology and Oncology–European Institute of Radiotherapy (ESTRO–EIR) report on 3D CT-based in-room image guidance systems: A practical and technical review and guide

Stine Korreman^a, Coen Rasch^b, Helen McNair^c, Dirk Verellen^d, Uwe Oelfke^e, Philippe Maingon^f, Ben Miinheer^b. Vincent Khoo^{c,g,*}

*Department of Radiation Oncology, The Finsen Centre, Rigshospitalet, Copenhagen, Denmark; *Department of Radiation Oncology, The Netherlands Cancer Institute/Antoni van Leeuwenhoek Hospital, Amsterdam, The Netherlands; *Department of Amical Oncology, Royal Marsden NHS Foundation Trust, Chelsea and Sutton, London, UK; *UZ Bussel, Oncologisch Centrum, Radiotherupie, Brussek, Belgium; *Department of Medical Physics in Radiation Oncology, Deutsches Krebforschungzentrum, Heddelberg, Germany *Popartment of Radiotherupie, Brussek, Belgium; *Department of Medical Physics in Radiation Oncology, Deutsches Krebforschungzentrum, Heddelberg, Germany *Department of Radiotherupie, Brussek, Belgium; *Department of Medical Physics in Radiation Oncology, Deutsches Krebforschungzentrum, Heddelberg, Germany *Department of Radiotherupie, Brussek, Belgium; *Department of Medical Physics in Radiation Oncology, Deutsches Krebforschungzentrum, Heddelberg, Germany *Department of Radiotherupie, Brussek, Belgium; *Department of Medical Physics in Radiation Oncology, Deutsches Krebforschungzentrum, Heddelberg, Germany *Department of Radiotherupie, Brussek, Belgium; *Department of Medical Physics in Radiation Oncology, Deutsches Krebforschungzentrum, Heddelberg, Germany *Department of Radiotherupie, Brussek, Belgium; *Department of Radiotherupie, Brussek, Brussek

Table 1

Factors for consideration in image acquisition and their relevance.

What field of view (FOV) length is available in the cranio-caudal direction?

Determines the length of scan available and possible solutions if longer scan lengths are required

What size is the reconstruction circle?

Determines the lateral FOV

Are filters required? - Which filters are available?

Involves time to select and insert, and affects image quality

Are filters interlocked?

If not, then risk of poor quality or unusable scans from incorrect filters selection

Can panel be positioned remotely? If so, does this the system come with an anti-collision system?

Will involve time to position if not remotely accessed

What are the available rotation speeds?

Determines the acquisition time

What are the possible angles of rotation?

Affects the flexibility of scanning; e.g. the possibility of performing half-scans for small regions, rotations through 180 degrees (underneath the patient) and using preset or flexible start and stop angles

How ergonomic is the operation?

One- or two-button operation, foot- or hand-control, several screens affects the ease of operation and the risk of aborted scans

Can the scan be stopped and restarted?

Will result in extra dose if the scan is interrupted inadvertently, and has to be started from the beginning Also allows the scan to be acquired with the patient in several breath holds.







Automatic Match

• Uses matching algorithm based on "Mutual Information" within the defined field of view

Manual Match

- Allows adjustments to be made using either mouse or keyboard
- User dependant
 - Respect the learning curve





- The Region of Interest Box
- Used for the automatic registration algorithm
- Defines the greyscale range (HU) that the algorithm will use for the solution
- The interface has additional options
 - ➤ Consider the "Stucture VOI" option
 - Margin added to this Structure VOI will help drive the MI algorithm
 - Intensity Range
 - > Be willing to adjust settings to ensure you are getting the most out of your system!
 - > Similar to Elekta, the anatomy included is very important



- Correctional shifts are displayed to the nearest 1mm
- Any automatic match *must* be reviewed by both the RTTs prior to treatment
- No machine can replace clinical judgement
- Know your volumes
 - Be aware of possibility of additional "planning volumes"

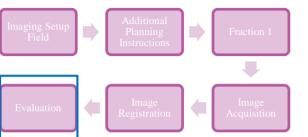






- How can we decrease inter observer variability?
 - Education of staff (encourage CPD, training packages, competency based assessment)
 - Protocolised imaging methods
 - Protocolised matching methods
 - Sequence of matching process
 - Automatic Match *must be followed by manual review* and adjustment
 - VOI and intensity levels set for each site and "locked" on Fx 1
 - Anatomy to include in VOI box

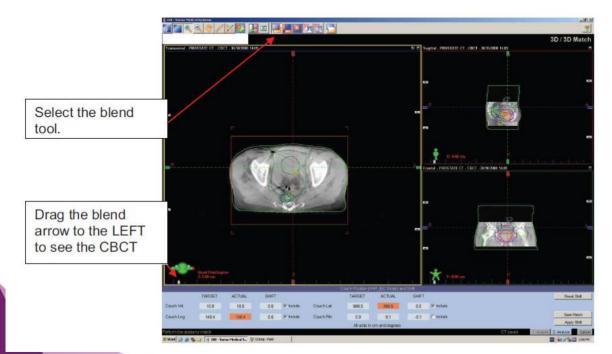




Processes available to assist in image evaluation



- Blending
 - Blending of the planned and acquired image
 - Colour or greyscale





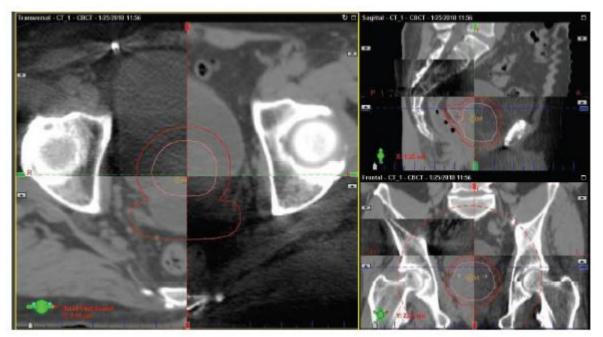




Processes available to assist in image evaluation



• Split screen



Don't forget to adjust the window level and move your views around



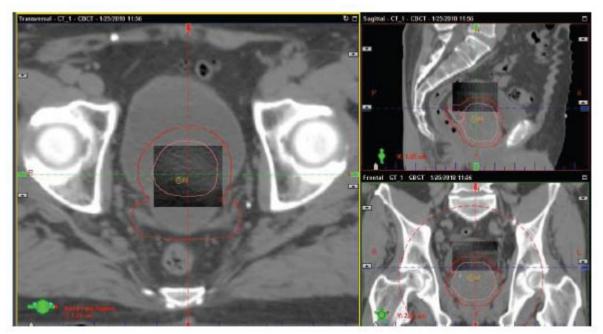




Processes available to assist in image evaluation



Moving window tool



Don't forget to adjust the window level and move your views around



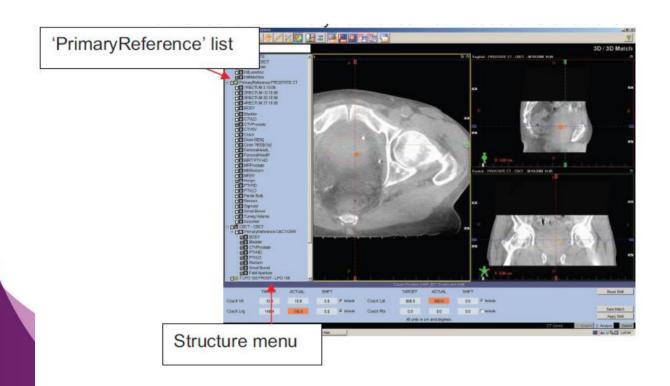




Processes available to assist in image evaluation



- Overlay Structure
 - Volumes that were contoured at the planning stage









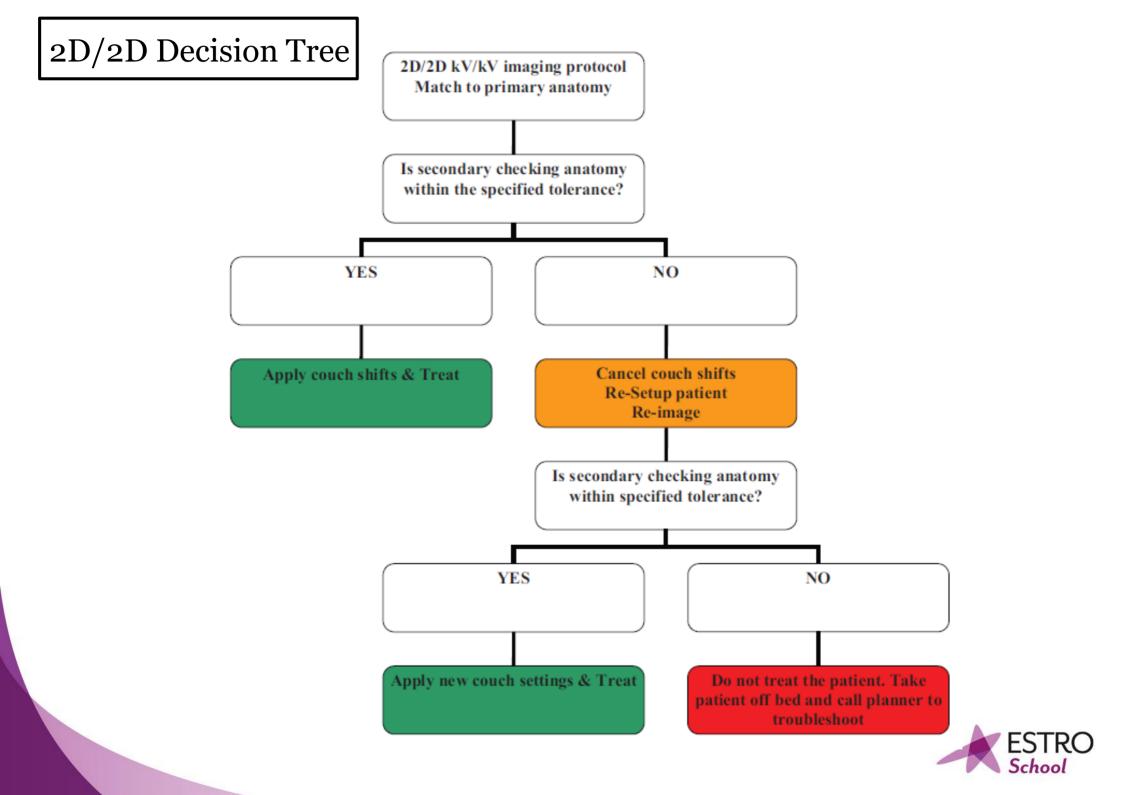
- The evaluation process must not be rushed
 - > Check that the shifts are *sensible*
- Both RTTs must confirm the match
- "If that were my mum..."
- It is better be check than treat the patient incorrectly
- IGRT is a team approach and if unsure there are always people to help
- Communicate!
 - > Journal, Alerts, annotation on the image





"the importance of this visual inspection cannot be over-emphasized and the user is encouraged to assess the accuracy of these automated registration tools" (Korreman et al., 2010)





CBCT imaging protocol Match to primary anatomy **CBCT** Decision Tree Is secondary checking anatomy within the specified tolerance? YES NO Is patient outline within Cancel couch shifts yellow tolerance rings? Re-Setup patient Re-image Is secondary checking anatomy within specified YES NO tolerance? Apply couch shifts & Treat Is patient outline within YES red tolerance rings? Is patient outline within YES Do not treat the patient. NO Take patient off bed and yellow tolerance rings? all planner to troubleshoot Apply couch shifts & Do not treat the patient. YES NO Take patient off bed and treat Add alert and schedule call planner to troubleshoot repeat CBCT Apply new couch settings & Treat Is patient outline within red tolerance rings? Apply couch shifts & Do not treat the patient. Take patient off bed and treat Add alert and schedule repeat CBCT call planner to troubleshoot





Site Specific Application







Radical CNS

 Examples of structures to outline on DRR for 2D/2D match

Site	Image type	Primary (red) and Confirming (yellow) matching anatomy	
Brain	KV & CBCT	2. Medial orbital rims 7. Superior orbital rims 8. Lateral skull wall (parietal region)	1. Pituitary fossa (volumend and transferred to reference image) 3. Frontal sinus 4. Base of skull in pituitary fossa region 5. Base of skull in Occipital bone region 6. Anterior cranial fossa

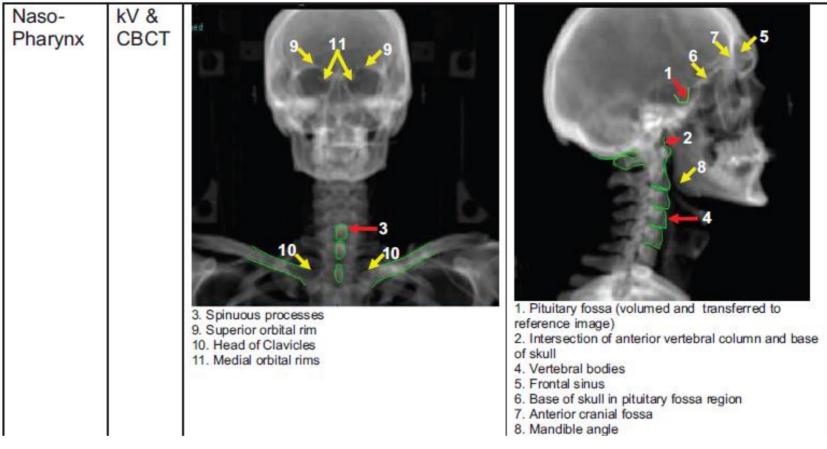






Head and Neck

 Examples of structures to outline on DRR for 2D/2D match

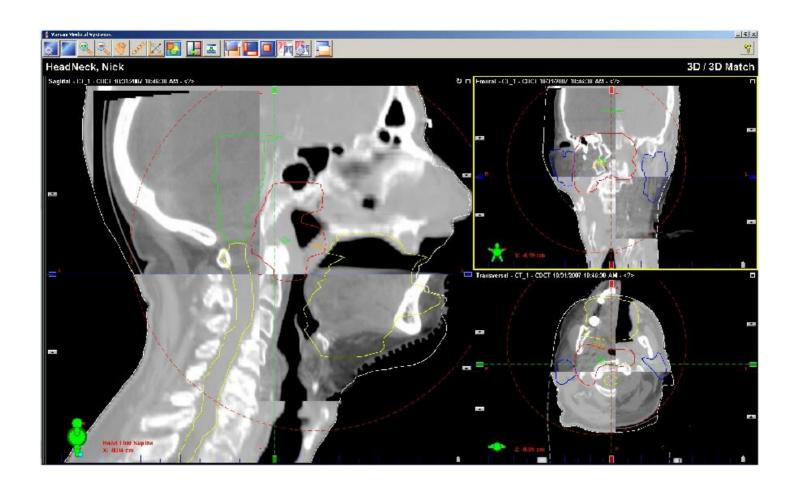








Head and Neck



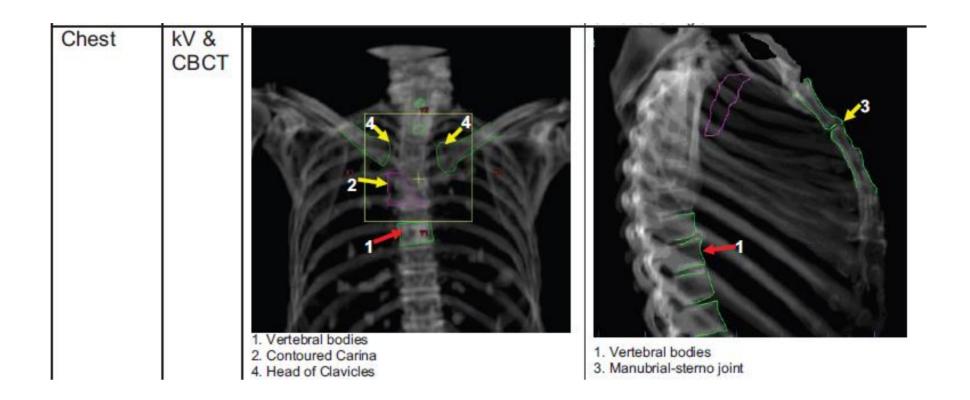






Thorax and Upper Abdomen

 Examples of structures to outline on DRR for 2D/2D match



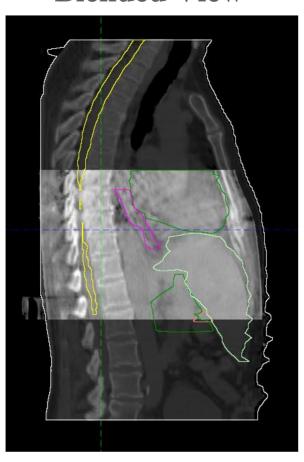




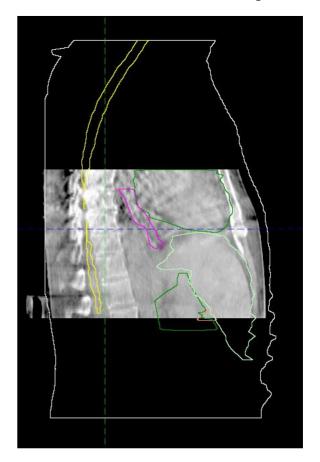


Thorax and Upper Abdomen

Blended View



Contour Overlay



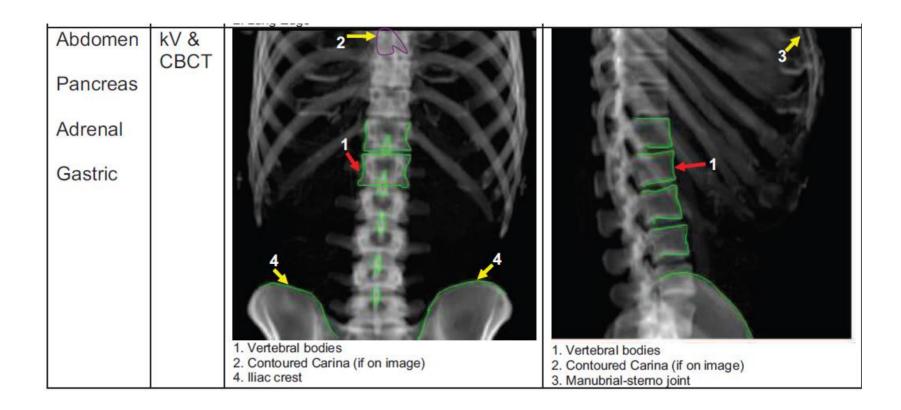






Abdomen

 Examples of structures to outline on DRR for 2D/2D match



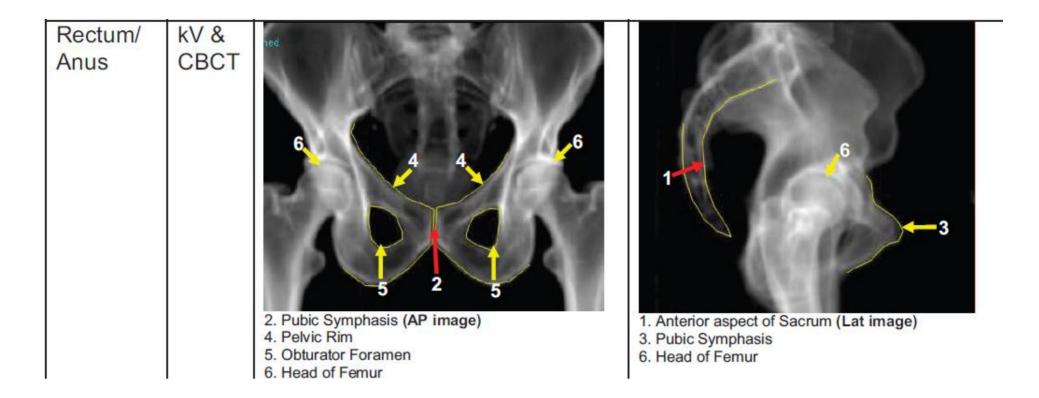






Rectum

 Examples of structures to outline on DRR for 2D/2D match





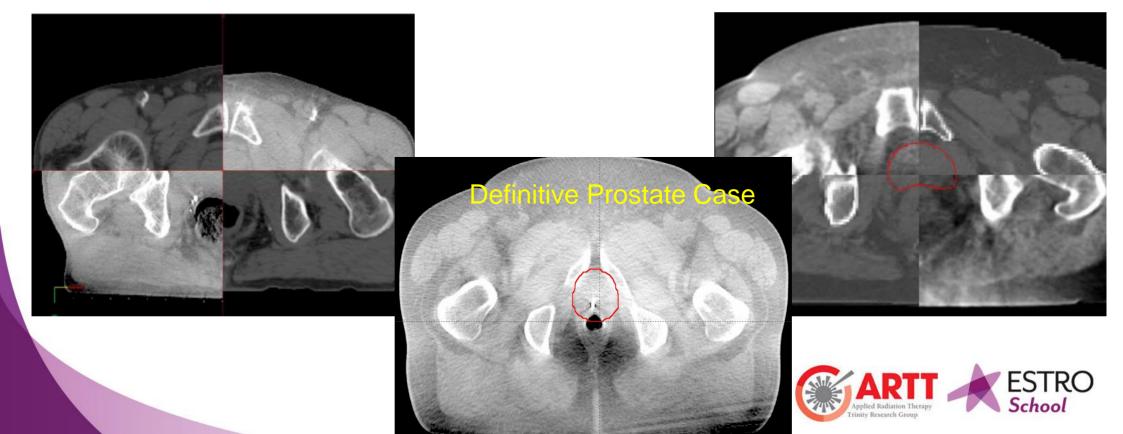




Prostate

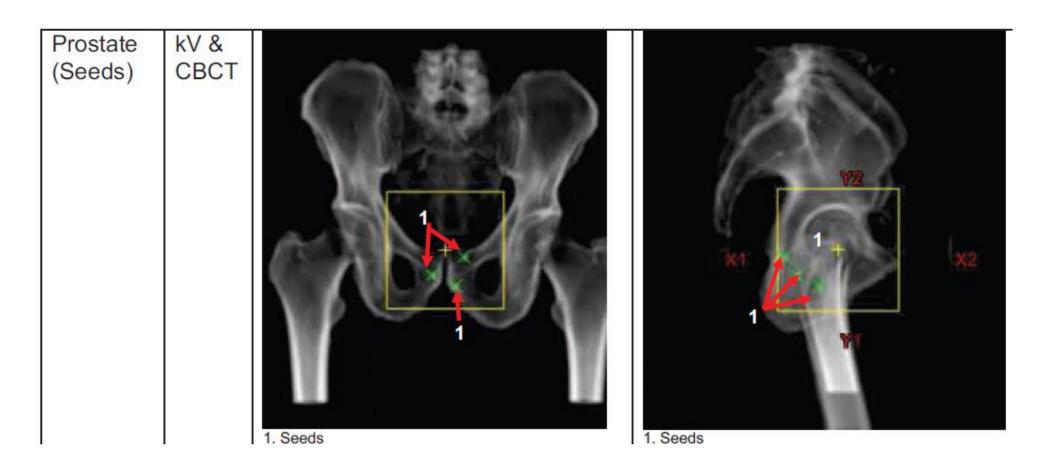
• Do **not** match to bones for definitive cases

Definitive Prostate	kV	All fractions except CBCT	Dellaman
(seeds)	CBCT	1,2,3,5,10,15,20,25,30,35,40	Daily moves
	•	•	
Definitive Prostate	CBCT	All fractions	Doily moyon
(soft tissue)			Daily moves



Prostate – 2D/2D fiducial match

Match points used for 2D/2D fiducial match

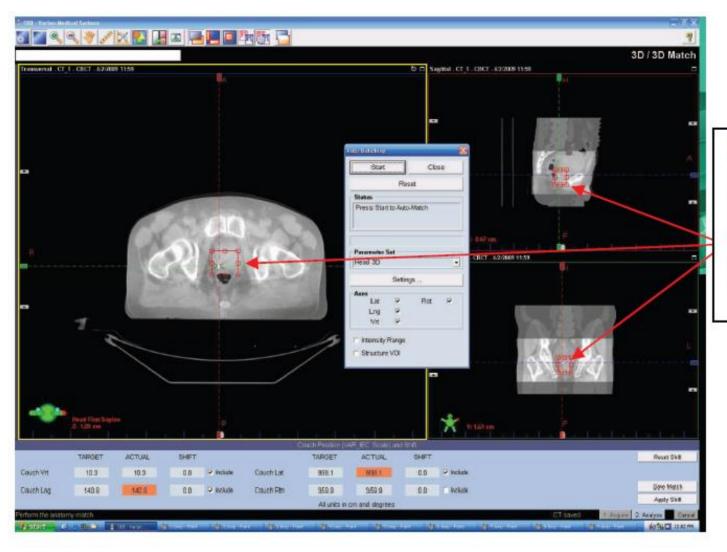








Prostate – CBCT fiducial match



Move the VOI box on all three views to cover the matching area.

Eg in this picture the fiducial markers

Always scroll through the entire length of the PTV and view in all 3 planes



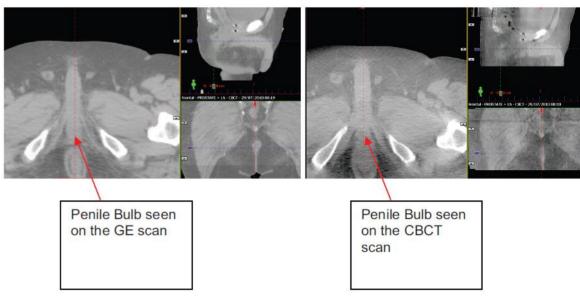
Prostate – Soft tissue

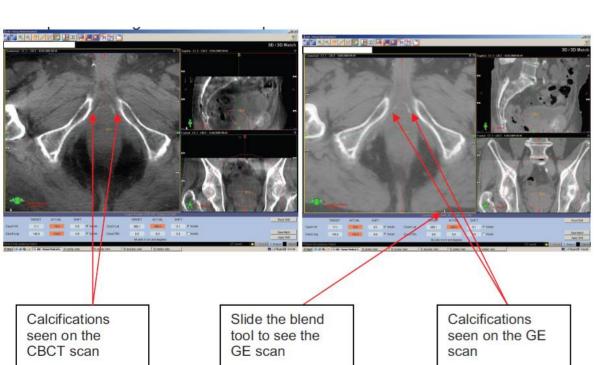
- Process for CBCT soft tissue match
 - ➤ *Manual confirmation* of match
- 1. Change window level to visualise rectum & superior prostate
- 2. Position superior CTV prostate contour to superior aspect of prostate at junction with bladder
- 3. Position posterior edge of CTV prostate structure (at mid prostate) to the anterior rectal wall
- 4. Check inferior CTV prostate structure to inferior edge of prostate, using penile bulb to assist
- 5. Position lateral edges of CTV prostate to pelvis muscles

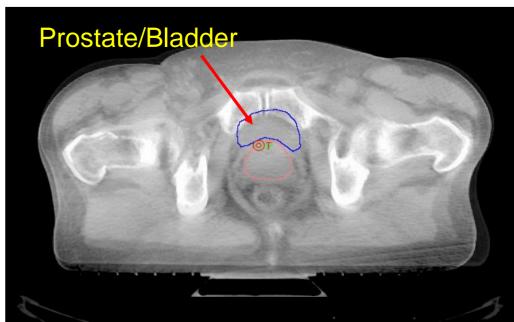


Prostate - Soft tissue









• Example of 2D anatomy to outline on the DRR

Prostate (Bone)	kV & CBCT	1. Pubic Symphasis	1. Pubic Symphasis
		2. Check any clips are covered with PTV for CBCT4. Pelvic Rim5. Obturator Foramen6. Head of Femur	Check any clips are covered with PTV for CBCT Anterior Sacrum Head of Femur







Troubleshooting

- These are all well suited and ideal cases
- What about when things aren't so clear?! *Troubleshoot*

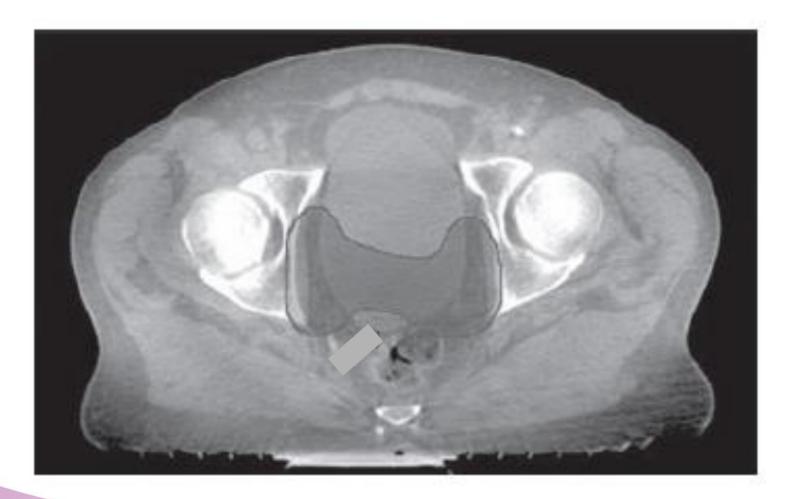






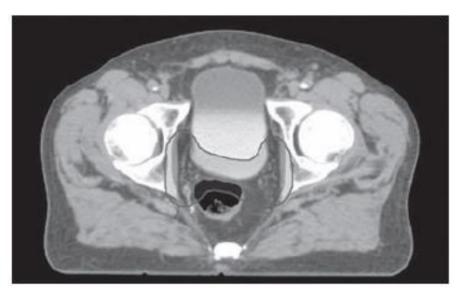


- Instructions match to bones
- Perfect match
- Isodose lines hug the PTV very nicely





 Have an anatomical understanding of exactly what the target is post surgery



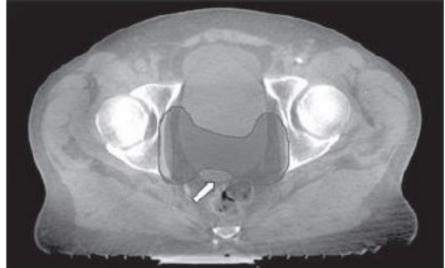


Fig. 4. CBCT of a patient undergoing radiotherapy following radical prostatectomy. Panel (a) shows the initial planning scan with the PTV displayed. Panel (b) shows a change in rectal volume resulting in the treated volume shifting outside the planning PTV (white arrow). CBCT, cone beam computed tomography; PVT, planning target volume.







Radiation Oncology-Original Article

Prostate bed motion may cause geographic miss in post-prostatectomy image-guided intensity-modulated radiotherapy

Linda J Bell^{1,2,*}, Jennifer Cox^{1,2}, Thomas Eade¹, Marianne Rinks^{1,†}, Andrew Kneebone¹

Article first published online: 9 JUL 2013

DOI: 10.1111/1754-9485.12089

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Journal of Medical Imaging and Radiation Oncology Volume 57, Issue 6, pages 725–732, December 2013

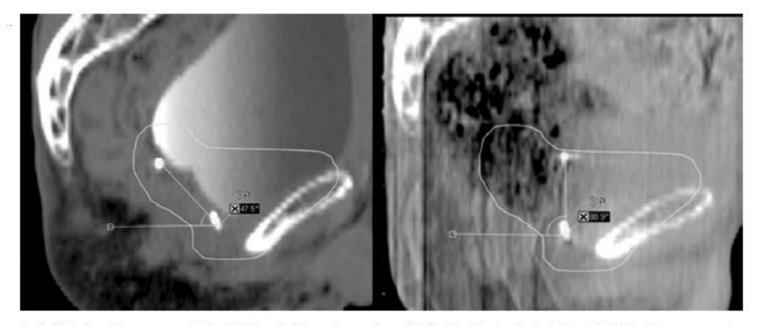
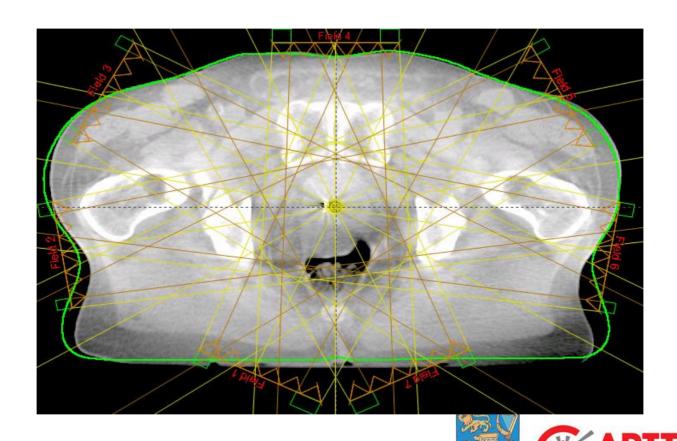


Fig. 1. Method used to measure prostate bed tilt. The angle between the superior and inferior clip relative to a horizontal line at the inferior clip was measured on the planning CT scan (left) and the cone beam CT scans (right). The angle-measuring tool in the Varian Offline Review® software was used to calculate this on the sagittal slice closest to midline of each scan where the clips could be visualised. The difference between the planning CT and cone beam CT angles was calculated. In this extreme case the angle on the planning CT (left) is 47.5° and that on the cone beam CT scan (right) is 88.9°. This is a difference of 41.4°. The FROGG-acceptable planning target volume expansion is delineated on these scans.



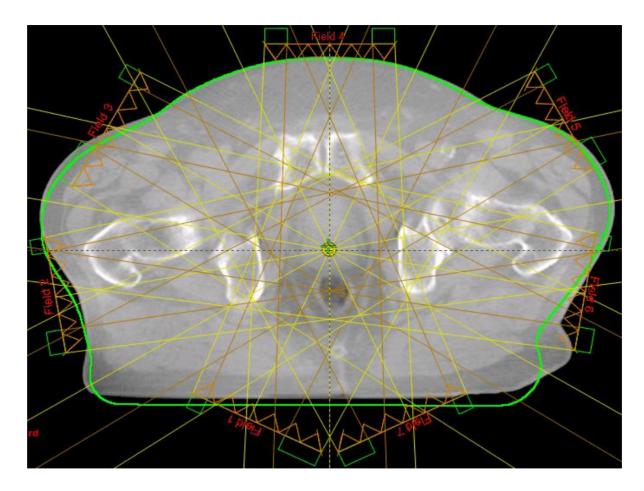
Definitive Prostate

- IMRT
- Daily online
- Match to implanted fiducials



Troubleshooting

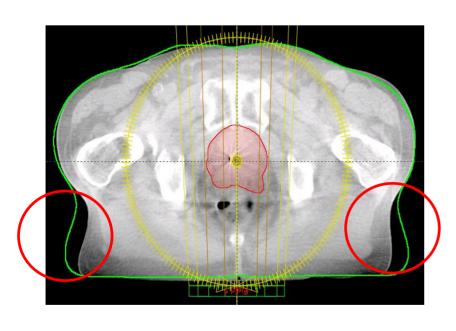
- Look beyond the target!
- Impact not on target *position*, but on target *dosimetry*

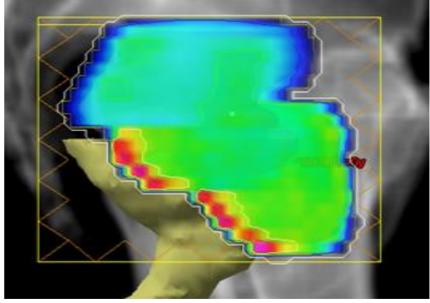


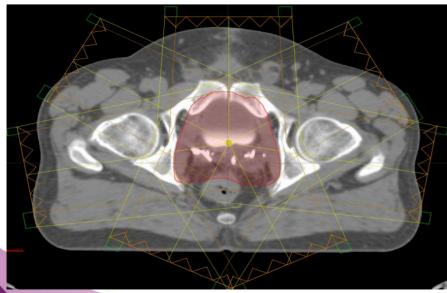


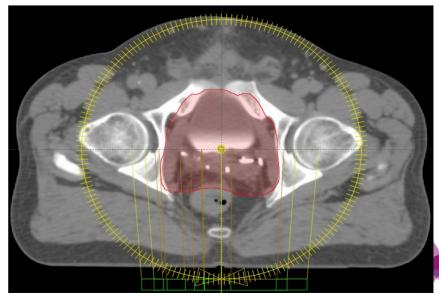
Troubleshooting

Integrate your planning knowledge – Clinical Intelligence!

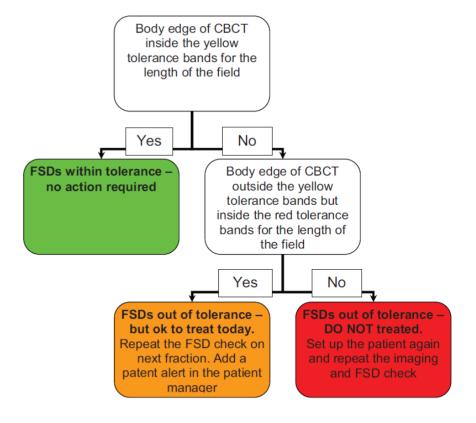








- What about when things aren't so clear?! *Troubleshoot*
 - Contour Variation
 - Weight Loss/Gain
 - Shoulder position
 - Neubauer et al 2012

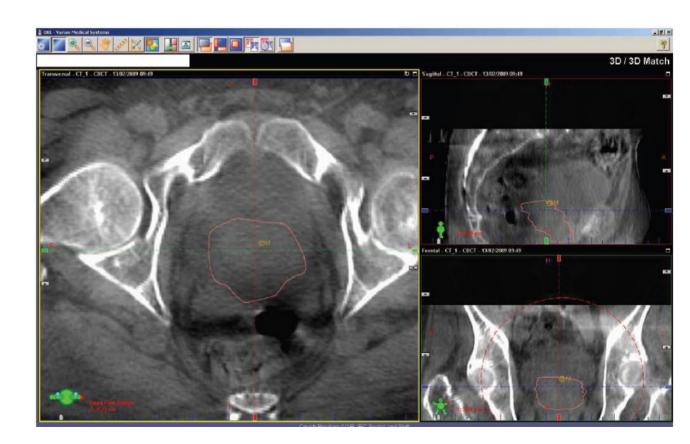






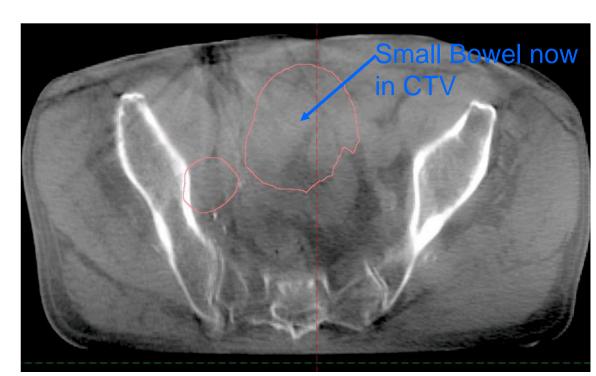


- What about when things aren't so clear?! *Troubleshoot*
 - Internal organ motion
 - Inter and intrafraction
 - o Gas

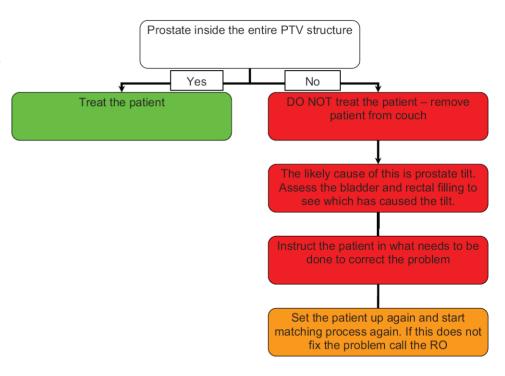


- What about when things aren't so clear?! *Troubleshoot*
 - Changes in bowel and bladder filling
 - Impact on target position and possibly dose
 - Impact on OAR dose

This is a bladder case, but also applicable to other sites (prostate bed)



- What about when things aren't so clear?! *Troubleshoot*
 - Displacement of CTV/PTV
 - Likely cause rotation or tilt
 - Motion of adjacent structures
 - Anatomical changes of target





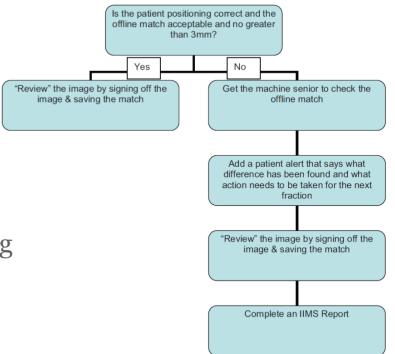




- What about when things aren't so clear?! *Troubleshoot*
 - Seed Migration
 - Poorly placed fiducials (SVs, Rectal wall etc)
 - > Need to guide the **system** here to ensure most appropriate match



- Online IGRT protocols should still include an offline review by an independent party
- This eliminates the time pressures of the machine
 - > RTT on machine
 - RTT in planning
 - > RO
 - Can also then feedback to patient
 - Patient education
 - Discuss at weekly MDT Audit Meeting









"The therapists are the front-runners for execution of the developed IGRT programs, and the quality of their performance will have a substantial impact on the success of IGRT" (AAPM Report 104)



Take Home Message!

- Use your "clinical intelligence"
 - Don't just automatch and hit apply to whatever the result is.
 - > **Think!** Does the match result make sense?
- Dosimetric Impact Thinking beyond the treatment unit
- Good idea to overlay the planned D95 or D100 isodose line on the CTV position
- Consider what is your target and what is the best surrogate for that
- Include the whole MDT



Acknowledgements

• Linda Bell, RTT and IGRT Specialist at Northern Sydney Cancer Centre



Brain





Jose Lopez, M.D., Ph.D Radiation Oncology University Hospital Virgen del Rocio Seville, Spain

Advanced skills in modern radiotherapy

Prague, Czech Republic –11-15 June 2016

Outline of Talk

- General pearls for Pediatric (CNS) tumors
- Protons
- Case report
- Discussion of current multidisciplinary (physician, phisyc and RTTs) management



Pearls

- The number one cause of death in children is accidents (44%), followed by cancer (10%).
- Of childhood cancers, leukemias are the most common followed by CNS neoplasms (~20%)
- Of pediatric CNS neoplasms, gliomas are most common (lowgrade astrocytomas ~35–50%, brainstem gliomas ~15%, malignant astrocytomas ~10%, optic pathway gliomas ~5%)



Inmovilization









Planning images

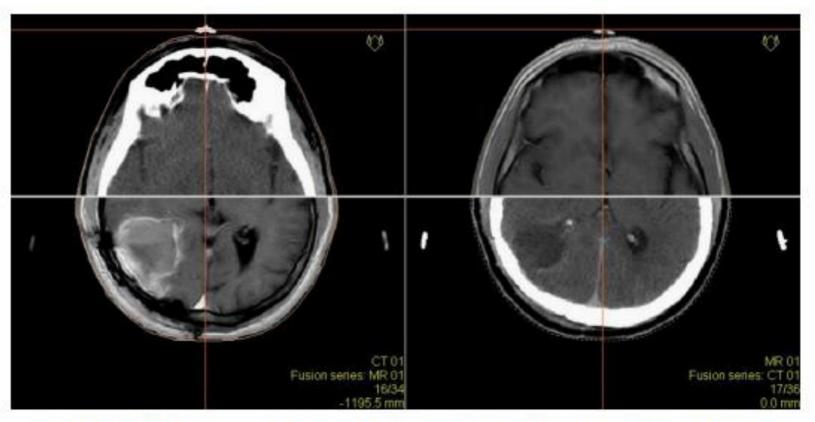


Fig. 2. Image registration of CT and MR image sets. Left image: top (CT), bottom (MR). Right image: top (MR), bottom (CT). The center of the middle fiducial marker pointed out on the left image is shown via registration on the right.



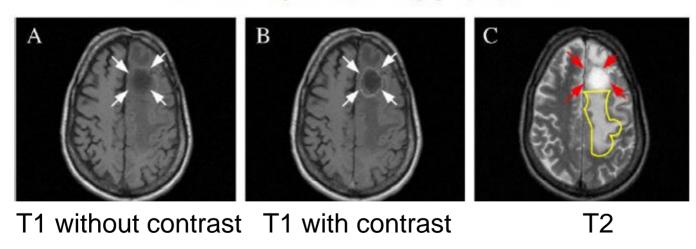


Fig. 4. Appearance of astrocytoma tumor on three sequences. (A) Isointense tumor (diffused) on T1-weighted image. (B) Isointense, peripherally enhanced homogeneous tumor on postcontrast T1-weighted image. (C) The tumor is seen as a homogeneous, hyperintense mass on T2-weighted image (in red) as is the edema, which, however, is less bright (in yellow).



T1 hyperintense

T1 isointense

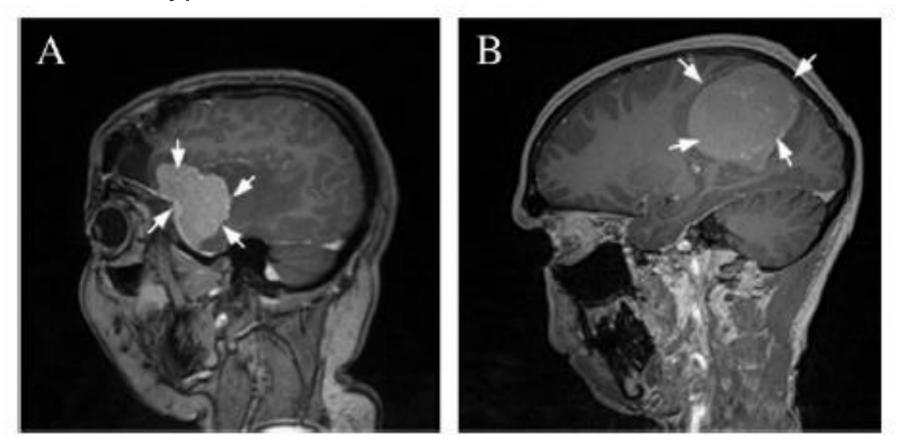


Fig. 2. Degree of enhancement of same tumor (meningioma) of different patients on postcontrast T1-weighted images. (A) Full enhancement of meningioma tumor (hyperintense signal). (B) No enhancement of meningioma tumor (isointense signal).



Homogeneous

Heterogeneous

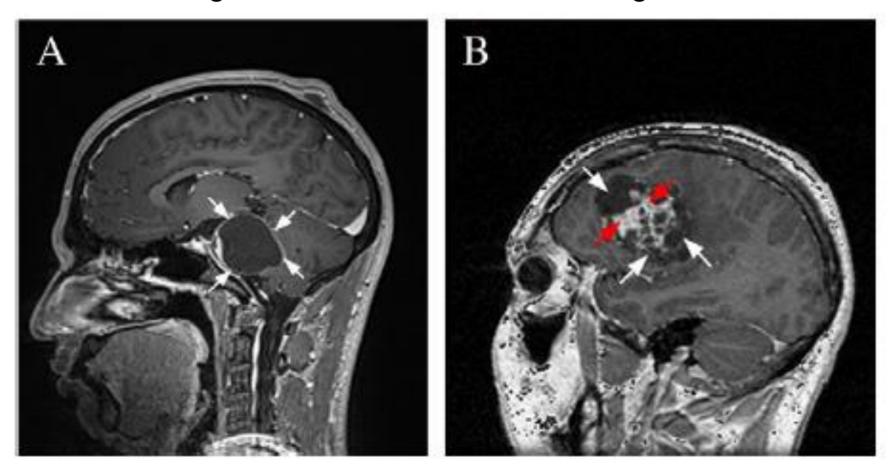


Fig. 3. Homogeneous and heterogeneous tumors. (A) Homogeneous astrocytoma tumor — hypointense signal, peripheral enhancement. (B) Heterogeneous glioma tumor with hypointense necrotic part (in red) and hyperintense-cystic components (in white).



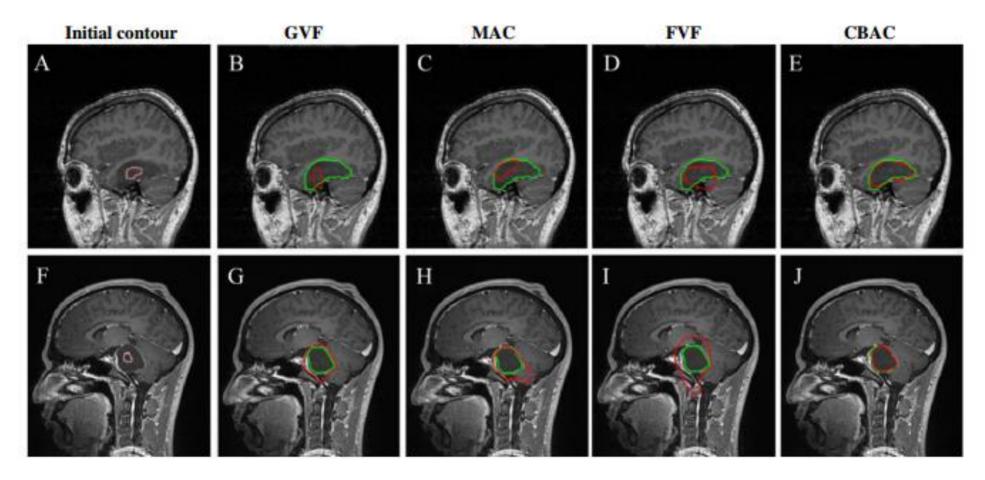
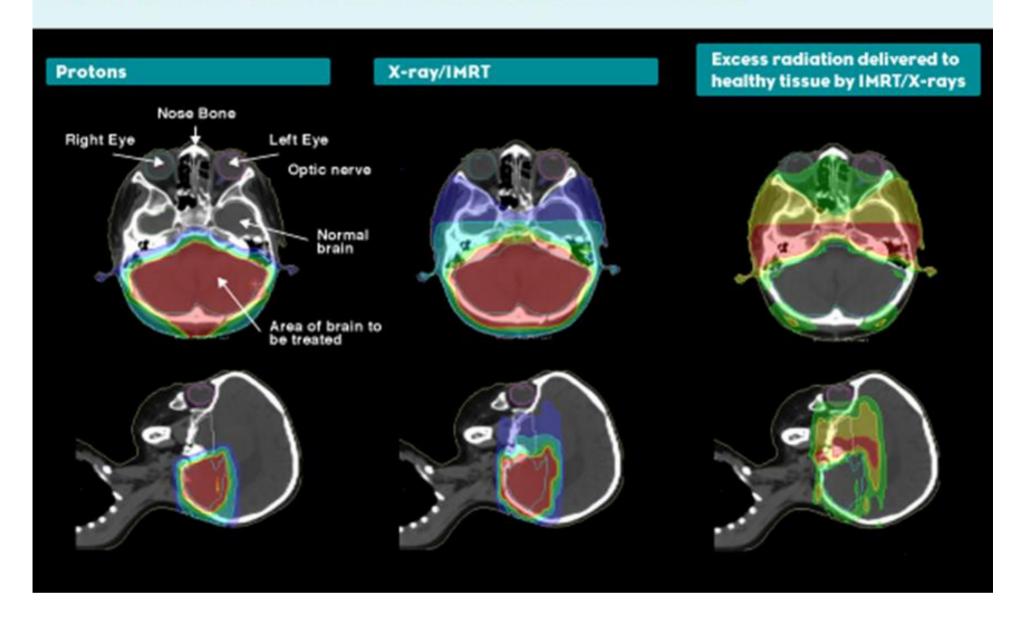


Fig. 10. Comparative segmentation results on postcontrast T1-weighted image. Green — ground truth marked by the radiologist; red — tumor boundary extracted by different methods. Row 1: tumor type, low-grade glioma; appearance, homogeneous tumor with isointense signal; the tumor shows no enhancement. Row 2: tumor type, astrocytoma; appearance, homogeneous tumor with hypointense signal; the tumor shows peripheral enhancement.



A Comparison of Radiation Treatment Plans for Pediatric Brain Cancer



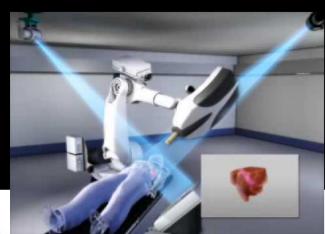


Technologies













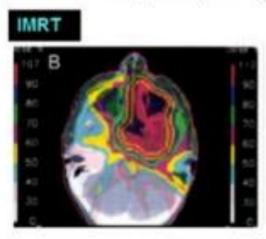
MV Cone-beam CT Approach

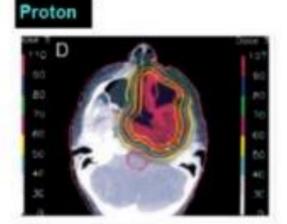


Proton Therapy

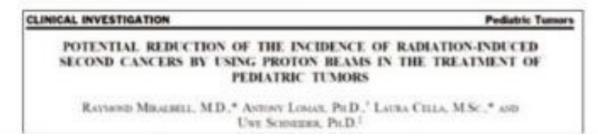
Claims:

- Opportunity to treat previously untreatable disease because of challenging geometry
 - Concave CTV/PTV partially surrounding a convex OAR





Second cancer reduction





Controversies of cost-effectiveness with new technologies, e.g. protons



is proton beam therapy for prostate cancer worth the cost?

February 20, 2013 By Durado Brooks, MD, MPH

Proton therapy popular and profitable

The lack of evidence has not slowed the rapid increase in the use of proton treatment for prostate cancer. One recent study documented a 67% increase in the number of cases of proton treatment for prostate cancer billed to Medicare between 2006 and 2009. This rate of growth is particularly noteworthy given the limited access to proton therapy: there are at present only 10 proton beam centers operating in the United States, and each center treats only a few hundred cancer patients each year.

- Cost of proton therapy nearly double compared to IMRT
- Benefit of proton therapy in prostate cancer is unproven
 - > Neither better tumor control nor lower toxicity
 - · A few studies suggest that toxicity rates might even be higher

pace? Financial incentives may be playing a role. Proton beam therapy for prostate cancer is reimbursed at a much higher rate than traditional radiation treatment for the same condition. Medicare pays about \$19,000 for a full dose of standard radiation therapy for prostate cancer, but it pays nearly double for proton therapy - more than \$32,000.



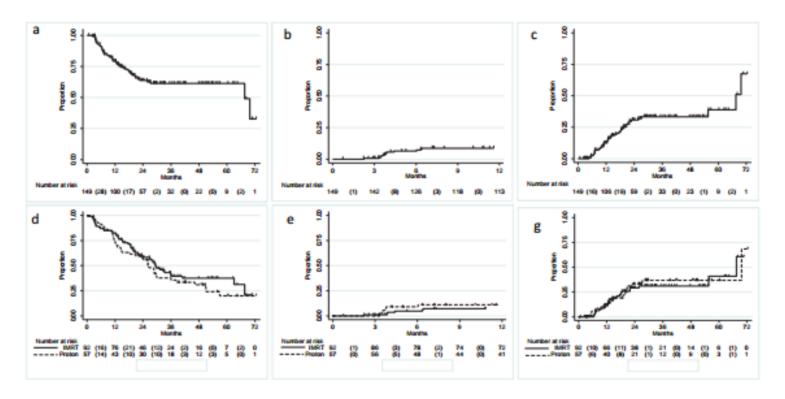
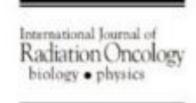


Figure 2. Time to treatment failure defined as 1) grade \geq 3 RP, 2) Local recurrence as first occurrence in randomised patients. Upper panels show the time to the development of (a) combined treatment failure, (b) grade \geq 3 RP, and (c) local recurrence as a whole group. Lower panels shox the comparison between IMRT (solid) vs. 3D-PBT (dashed) in time to the development of (d) combined treatment failure, (e) grade \geq 3 RP, and (g) local recurrence.



Lung cancer
Photons vs protons
Phase III
MDA





www.redjournal.org

Clinical Investigation: Late Effect

Incidence of Second Malignancies Among Patients Treated With Proton Versus Photon Radiation

Christine S. Chung, MD, MPH, * Torunn I. Yock, MD, MCh, Kerrie Nelson, PhD, Yang Xu, MS, Nancy L. Keating, MD, MPH, * and Nancy J. Tarbell, MD

Vol. 87, No. 1, pp. 46-52, 2013

- Conflicting hypotheses:
 - whether proton radiation has less risk than photon therapy
 - scattering with photons vs neutron contamination with protons
- 558 proton pts treated from 1973 to 2001 Harvard cyclotron vs 558 matched photon pts from SEER
- Second Ca: 29 protons (5.2%) and 42 Photons (7.5%)
- No evidence for or against (adjusted for age at treatment, sex, site, year diagnosed)



A systematic literature review of the clinical and cost-effectiveness of hadron therapy in cancer

Mark Lodge^{a,*}, Madelon Pijls-Johannesma^b, Lisa Stirk^c, Alastair J. Munro^d, Dirk De Ruysscher^{b,e}. Tom Jefferson^a

*Cochrone Cancer Network, Oxford, UK, *MAASTRO Clinic, Maastricht, The Netherlands, *Centre for Reviews & Dissemination, University of York, UK, *University of Dundee, Scotland, UK, *University Hospital Maastricht, GROW, MAASTRO Clinic, Maastricht, The Netherlands

Table 1
Results literature review in comparison with conventional therapy classified by tumour site

Tumour site	Protons		lons		
	n studies/N	Result	n studies/N	Result	
Head and neck	2/62	No firm conclusions	2/65	Similar to protons	
ACC (locally advanced)	-		1/29	Superior	
Prostate cancer	3/1751	Similar	4/201	No firm conclusions	
Ocular tumours	10/7708	Superior	2/1343	Similar to protons	
Gastro-intestinal cancer	5/369	No firm conclusions	2/73	No firm conclusions	
Lung cancer (non-small cell)	3/156	No firm conclusions	3/205	Similar to SRT	
CN5°	10/839	Similar	3/405	Similar to protons	
Chordomas of skull base	3/302	Superior	2/107	Similar to protons	
Sarcoma's	1/47	No firm conclusions	1/57	No firm conclusions	
Pelvic tumours	3/80	No firm conclusions	2/49	No firm conclusions	

Abbreviations: N, number of patients; ACC, adenoid cystic carcinomas; SRT, stereotactic radiotherapy.

- Brada et al. (JCO 2008) concluded that there is insufficient evidence at the present to recommend the use of proton therapy in any disease sites
- Reviewers / Authors have different views as to what constitutes evidence



^{*} CN5, central nerve system tumours; inclusive skull base, spinal cord chondroma and chondrosarcomas.

Radiotherapy and Oncology 103 (2012) 8-11



Contents lists available at SciVerse ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



Systematic review

An evidence based review of proton beam therapy: The report of ASTRO's emerging technology committee

Aaron M. Allen a.*, Todd Pawlicki b, Lei Dong c, Eugene Fourkal d, Mark Buyyounouski d, Keith Cengel c, John Plastaras c, Mary K. Bucci c, Torunn I, Yock c, Luisa Bonilla d, Robert Price d, Eleanor E, Harris G, Andre A, Konski b

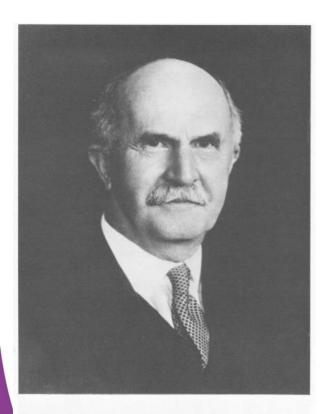
- In pediatric CNS malignancies PBT appears superior to photon approaches but more data is needed.
- In large ocular melanomas and chordomas, we believe that there is evidence for a benefit of PBT over photon approaches.
- PBT is an important new technology in radiotherapy
 - Current evidence provides a limited indication for PBT
 - More robust prospective clinical trials are needed to determine the appropriate clinical setting for PBT

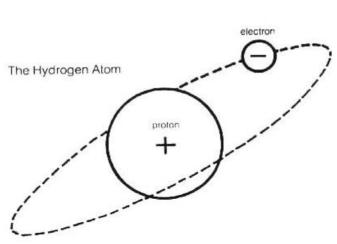


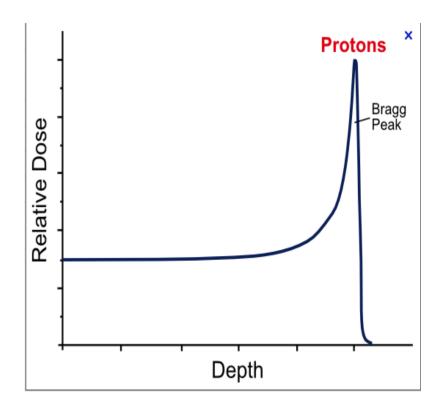




BRAGG PEAK

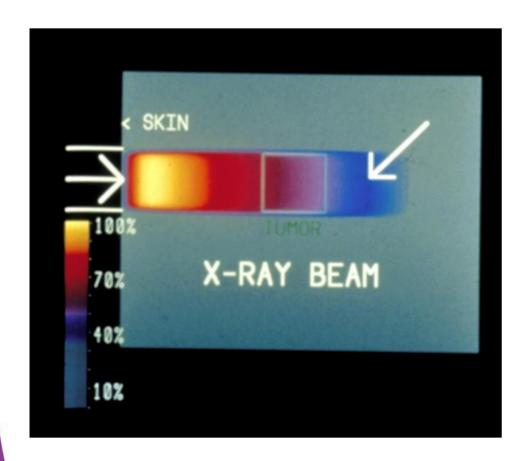


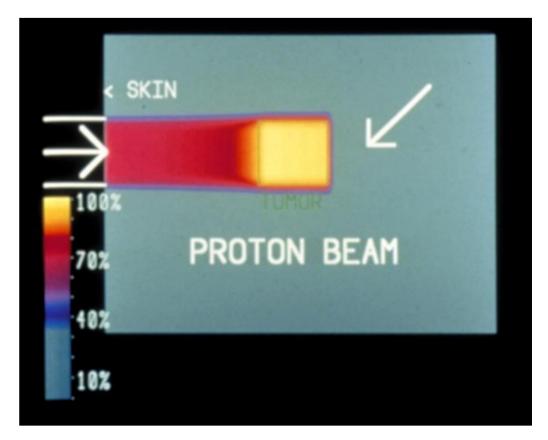




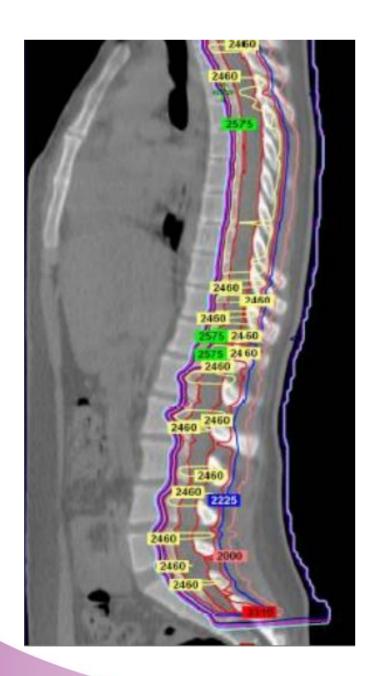


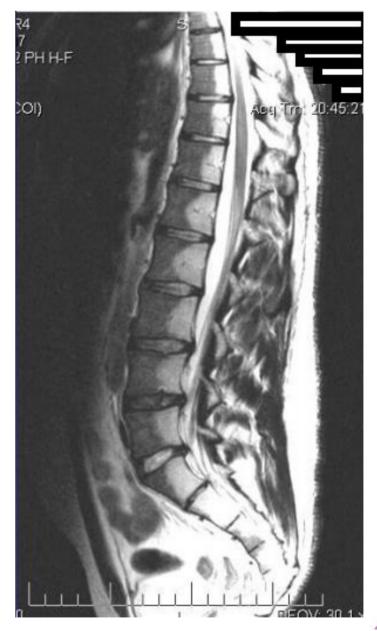












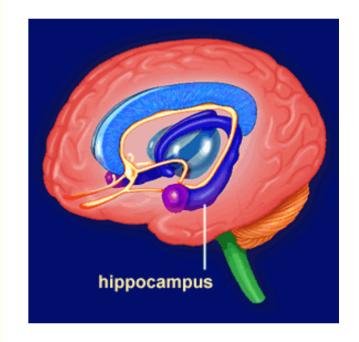


PTC

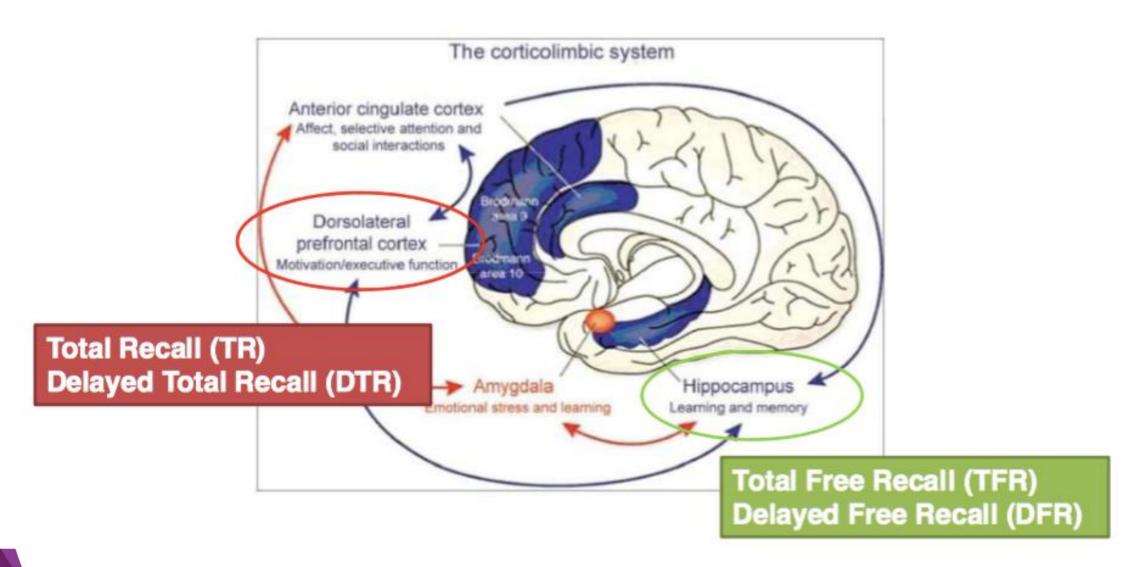


Association between hippocampal dosimetry and impairment in Wechsler Memory Scale-III Word Lists Delayed Recall at 18 months

Dosimetry	Dosimetric	No	Impairment*	p
	cut point	impairment	:	value
Bilateral hipp	ocampi			
Maximum	≤24.7 G y	66.7%	33.3%	0.500
	>24.7 Gy	55.6%	44.4%	
D30%	≤8.2 Gy	77.8%	22.2%	0.167
	>8.2 Gy	44.4%	55.6%	
D40%	≤7.3 G y	88.9%	11.1%	0.025
	>7.3 G y	33.3%	66.7%	
D50%	≤3.8 G y	66.7%	33.3%	0.500
	>3.8 Gy	55.6%	44.4%	
D80%	≤o.5 Gy	55.6%	44.4%	0.500
	>0.5 Gy	66.7%	33.3%	
D100%	≤o.o Gy	76.9%	23.1%	0.047
	>o.o Gy	20.0%	80.0%	
Left hippoca	mpus			
Maximum	≤15.0 Gy	55.6%	44.4%	0.500
	>15.0 Gy	66.7%	33.3%	









SPANISH LUNG GROUP 2017

	FIRST TIME	TOTAL FREE RECALL (TFR)	FREE RECALL (TR)	DELAYED FREE RECAL (DFR)	DELAYED T. RECALL (DTR)	
BASAL-3						
PCI	2 (6,7%)	4 (13,3%)	7 (23,3%)	8 (26,7%)	8 (26,7%)	
Hippocampal sparing	4 (13,3%)	2 (6,7%)	3 (10%)	1 (3,3%)	5 (16,7%)	
	NS	NS	NS	0,01 RR 8 [1,06- 60,08]	NS	
BASAL-6						
PCI	11 (40,7%)	9 (33,3%)	14 (51,9%)	13 (48,1%)	14 (51,9%)	
Hippocampal sparing	3 (14,3%)	1(4,8%)	3 (14,3%)	1 (4,8%)	5 (23,8%)	
	0,06 RR 2,8 [0,9- 8,9]	0,01 RR 7 [0,9- 50,9]	0,01 RR 3,6 [1,19- 11,0]	0,001 RR 10 [1,4- 71,23]	0,07 RR 2,1 [0,9- 5,08]	

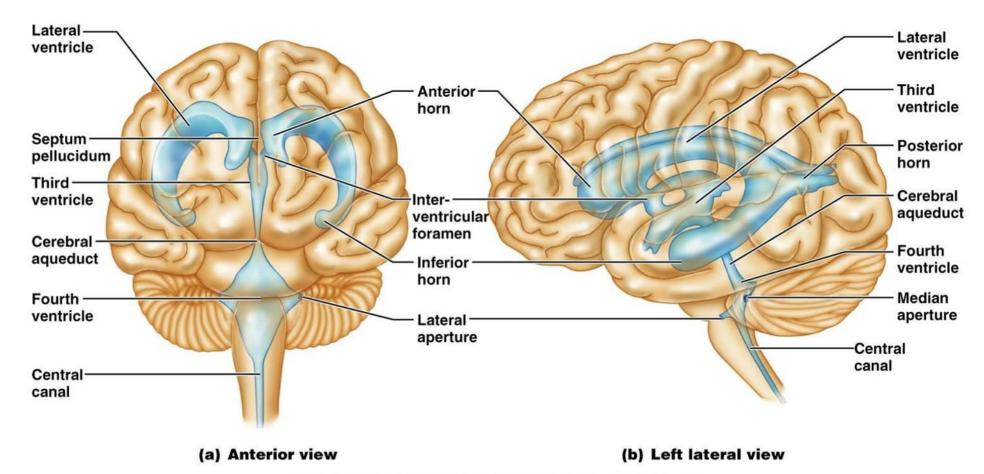


Case 1: patient with teratoid rhabdoid tumor

- A 19-month-old female infant was referred because of headache and weakness
- Magnetic resonance imaging revealed a mass that occupied the fourth ventricle



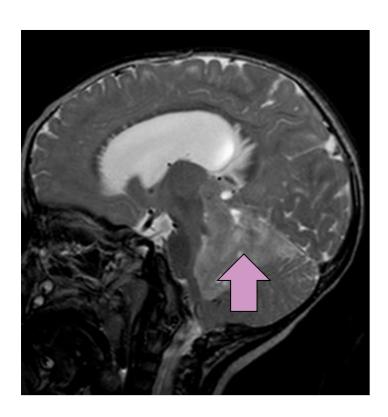


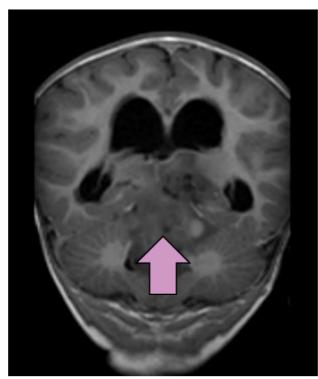


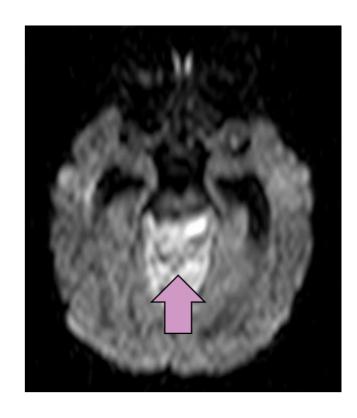
Copyright © 2006 Pearson Education, Inc., publishing as Benjamin Cummings.



Mass at the fourth ventricle

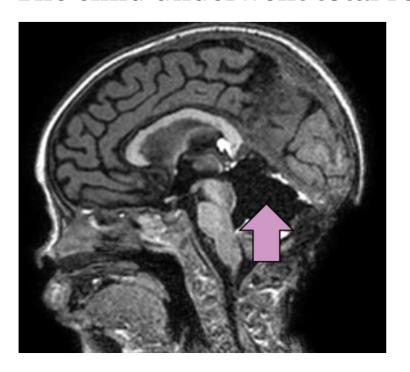


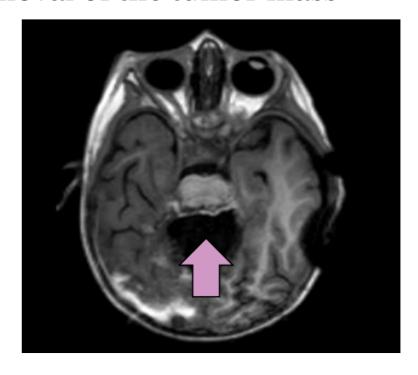






• The child underwent total removal of the tumor mass





 Pathological findings showed an atypical teratoid/rhabdoid tumor

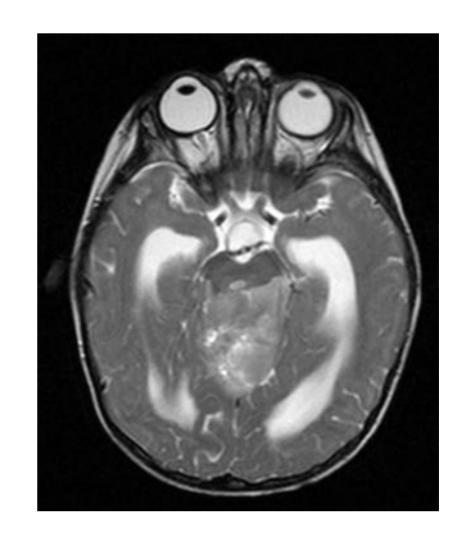


- Diagnosis
- Atypical teratoid/rhabdoid tumor
- Treatment
- Chemotherapy + Surgery + Radiation Therapy
- Radiation Therapy Dose Prescription:
- PTV (surgical bed + 5mm margin): 54 Gy at 2 Gy/fraction



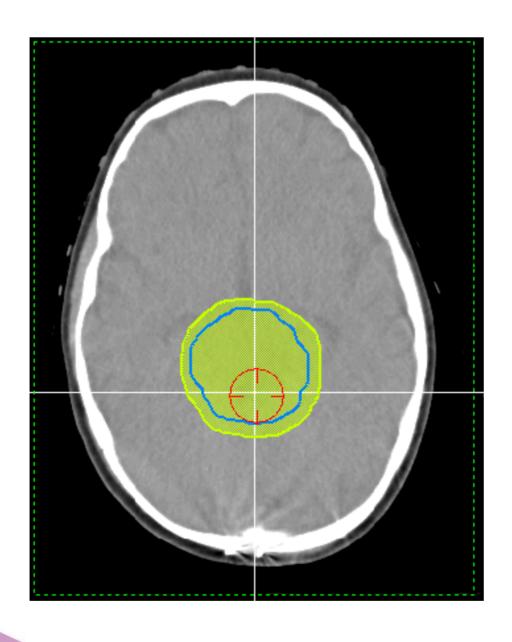
Organ at risk

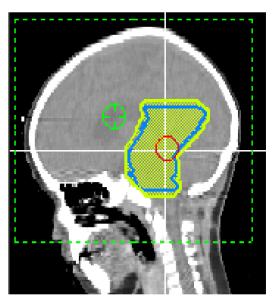
Whole brain
Braim stem
Chiasm
Pituitary
Eyes
Crystalline lens
Nerve optic

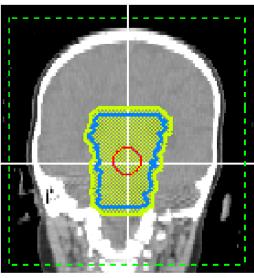




PTV (surgical bed + 5 mm margin)

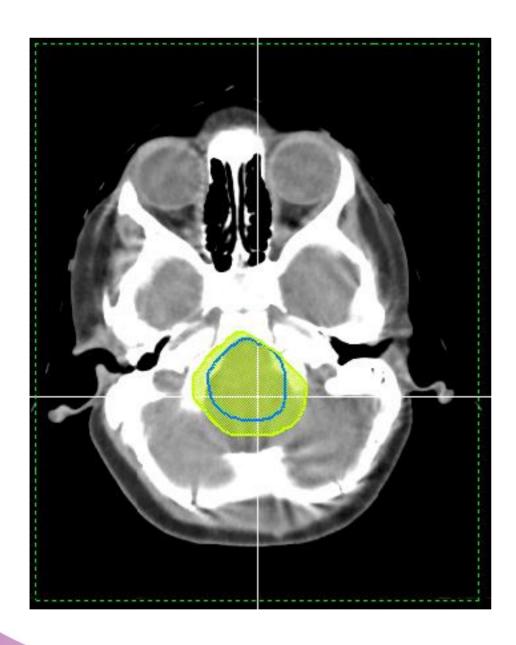


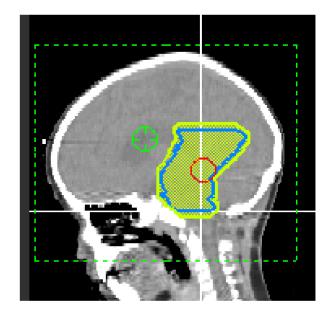


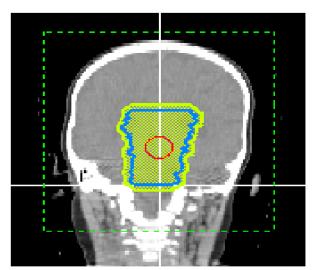




PTV(yellow)









Take home message

Inmovilization is crucial to reduce toxicity

• The addition of MRI gives vastly superior softtissue visualization

• The radiation technique (IMRT, Tomotherapy, Protons, Cyberknyfe) should be individualised for each patient





Questions:

- Preparation (thermoplastic mask)
- Positioning
- Organ at risk contouring
- Set-Up
- Verification
- Radiation technique







Case reports: **Brain** a physicist's perspective



Mirjana Josipovic

Dept. of Oncology, Rigshospitalet & Niels Bohr Institute, University of Copenhagen

Denmark

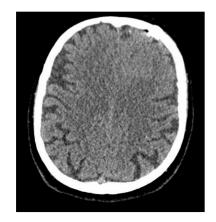
Advanced skills in modern radiotherapy
June 2017



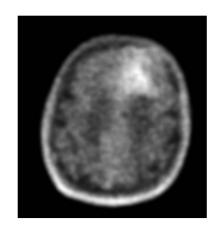
Imaging for brain RT planning

Imaging immobilised patient in the treatment position

CT scan

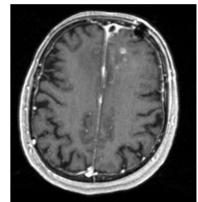


FET PET scan

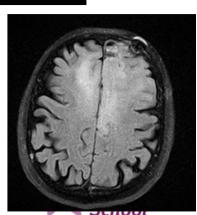


Thin scan slices ~1 mm

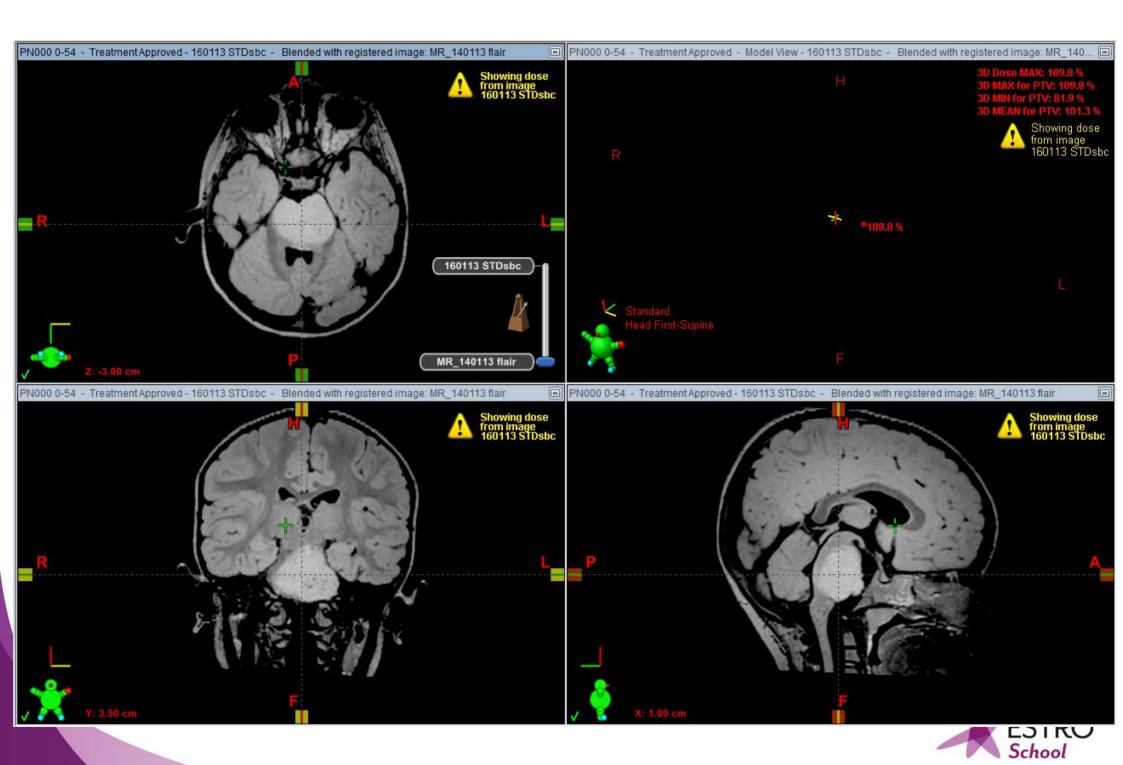
MR scan (T1,T2,FLAIR)

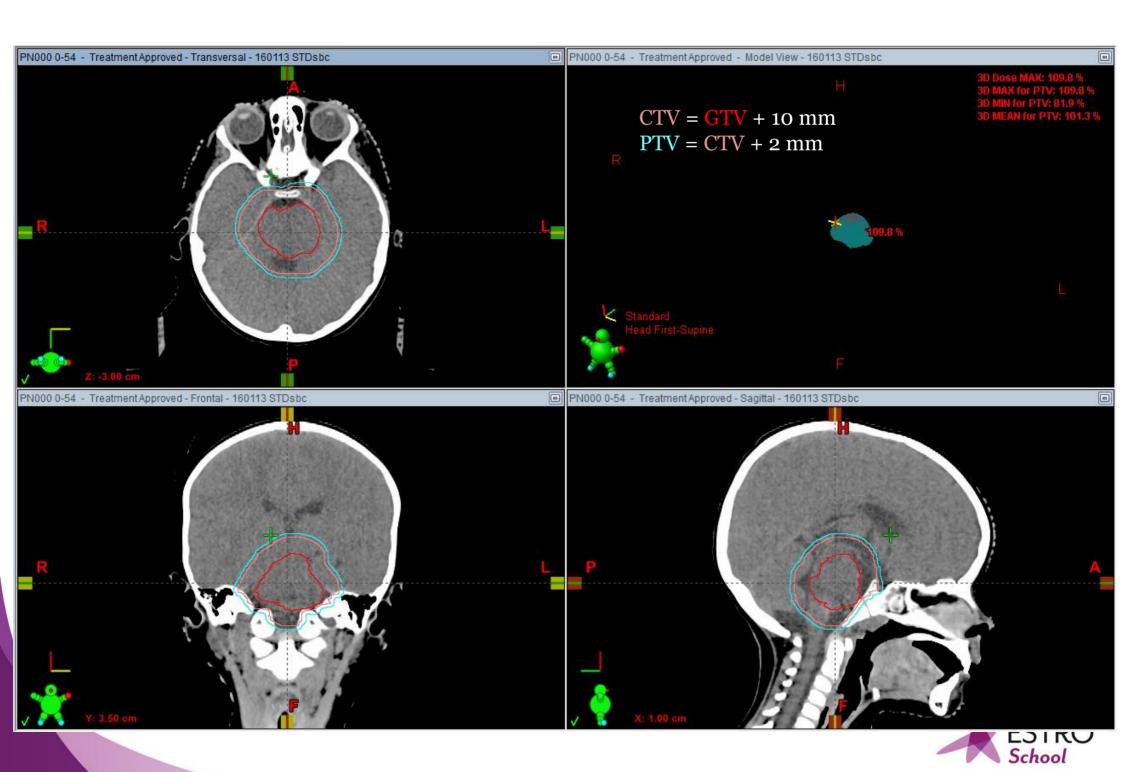






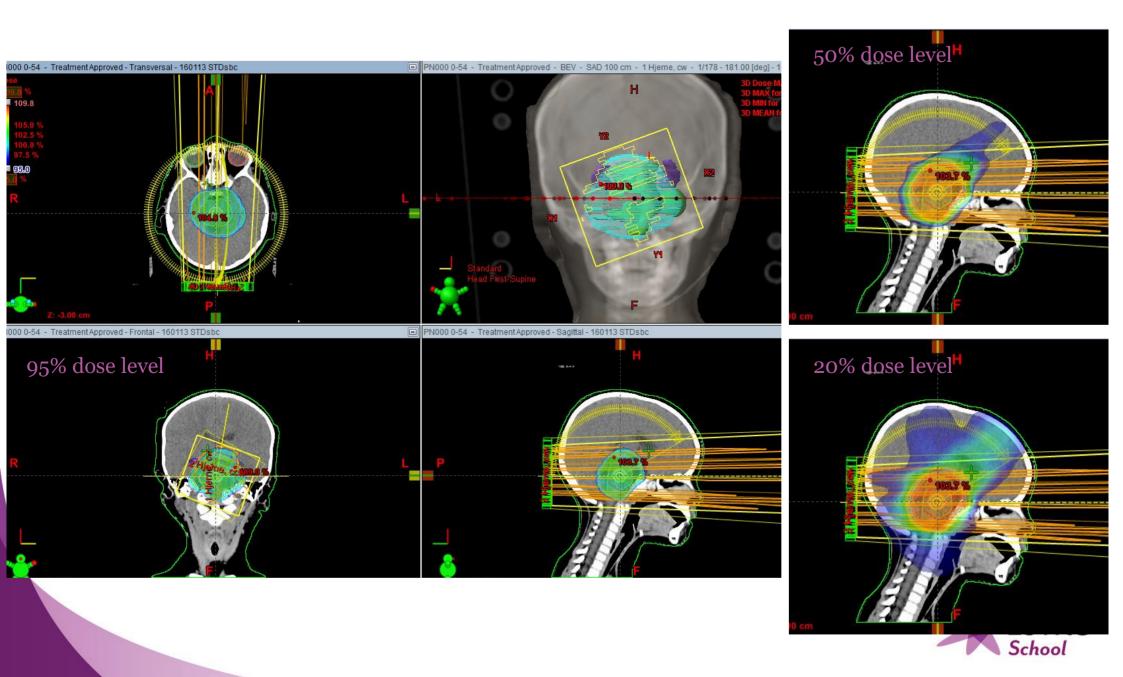








VMAT plan – 2 arcs



Radiotherapy

Radiotherapy techniques

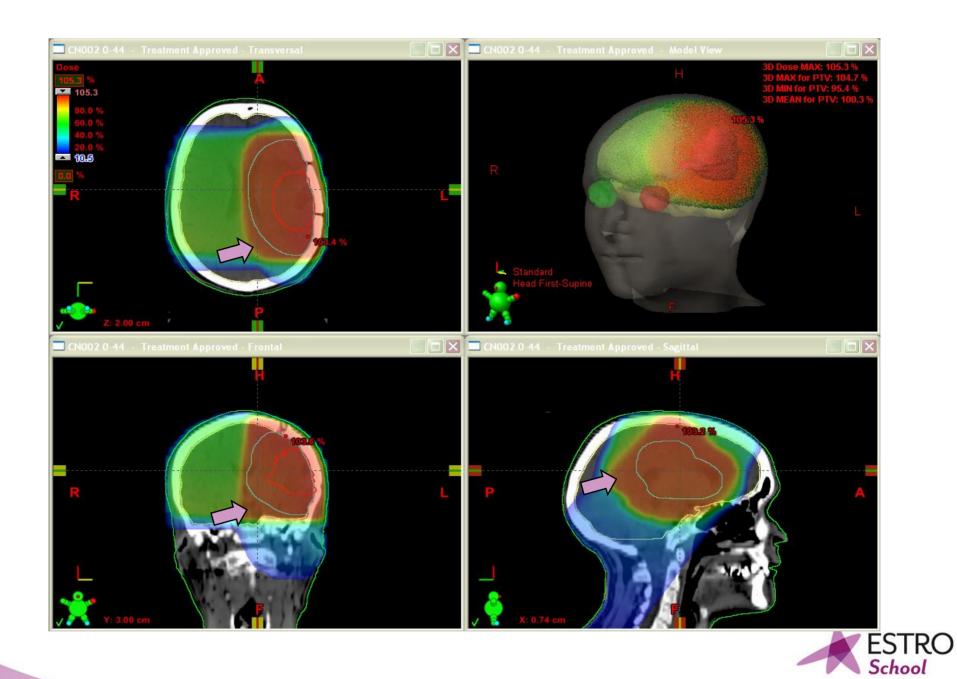
- 3DC
- IMRT
- VMAT
- Proton therapy

Fractionation schemes (Rigshospitalet, CPH)

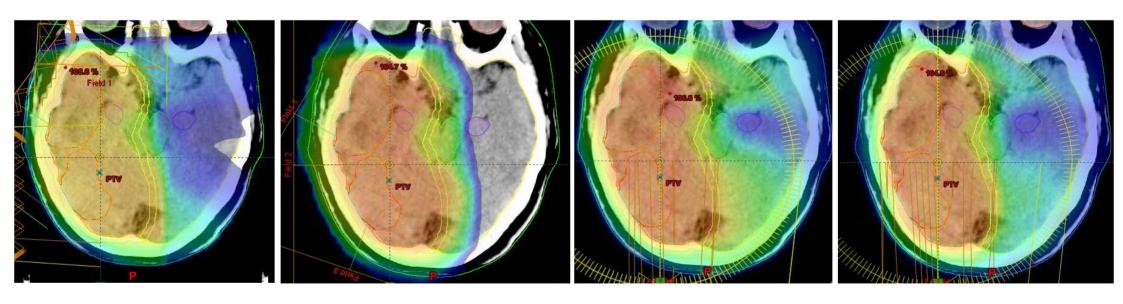
- 2 Gy x 30
- 1.8 Gy x 30 (if brainstem is involved)
- 18 Gy x 1 (very small targets, stereotactic RT)
 - Prescribed as minimum dose to target



3DC plan



IMRT vs. protons vs. VMAT



IMRT

IMPT

protons

VMAT (co-planar)

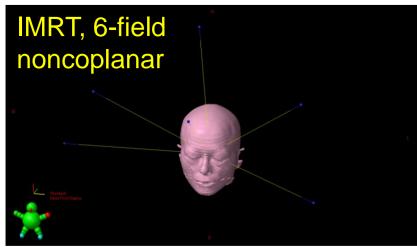
VMAT (non co-planar)

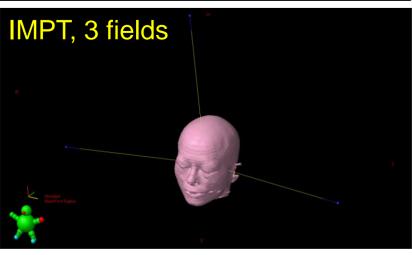
worst plan conformity

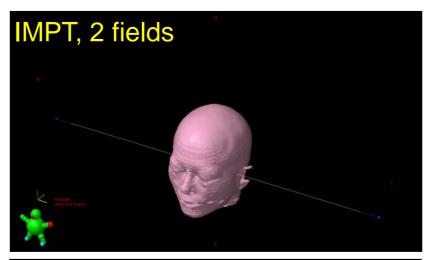
best plan conformity

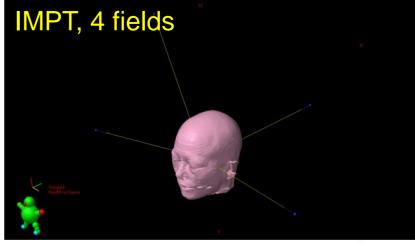


Many geometrical possibilities



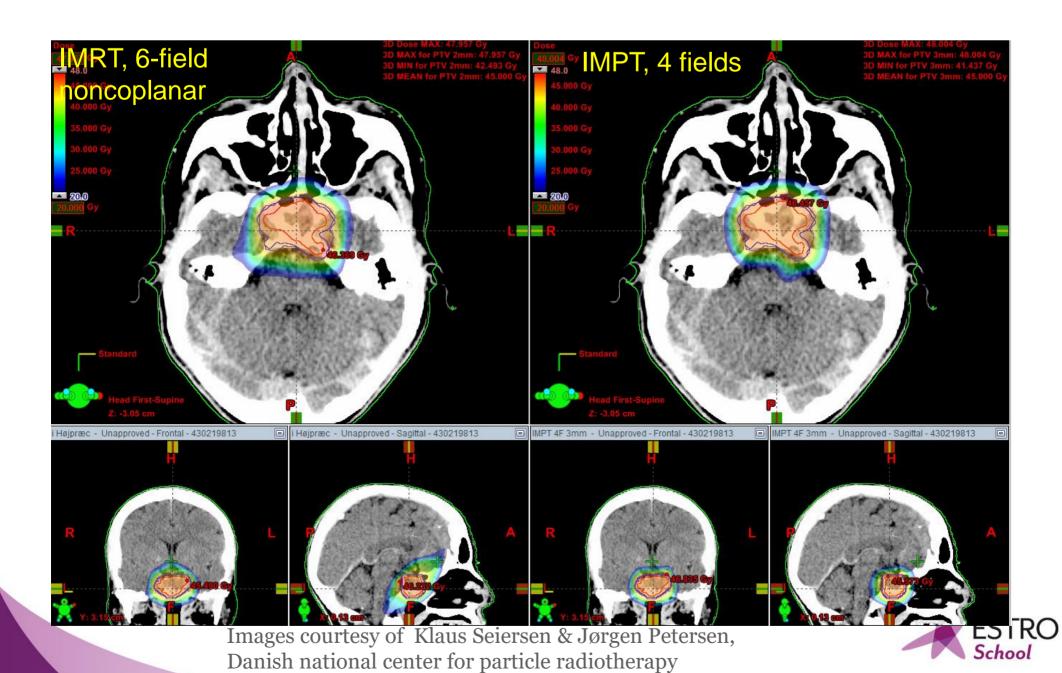


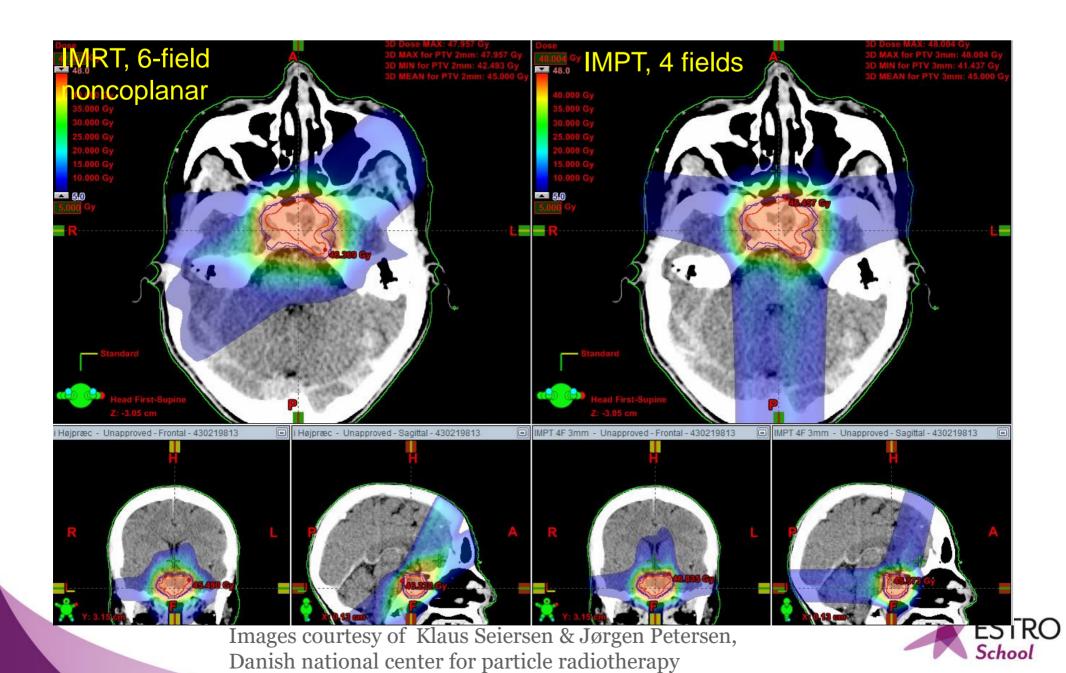




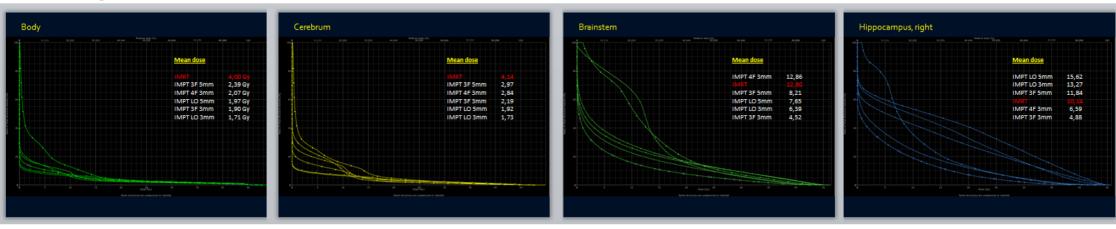


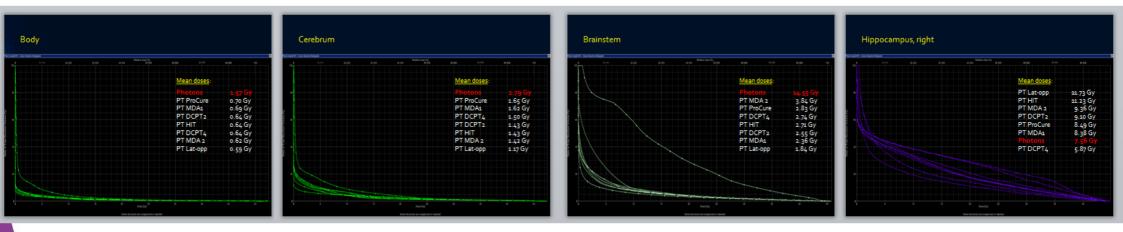
Images courtesy of Klaus Seiersen & Jørgen Petersen, Danish national center for particle radiotherapy





RED = photon **IMRT**







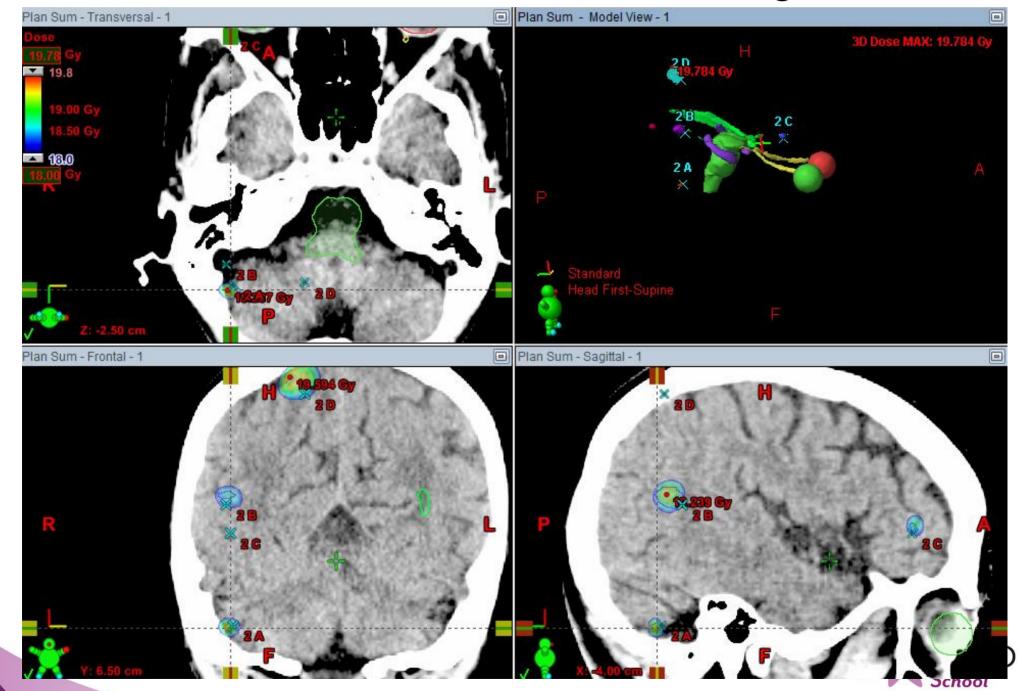
- OAR sparing depends on choice of treatment fields/angles
- Which organ to prioritise?
 - Doctor's order = "dose as low as possible"
- Integral dose is always lower with protons
 - Risk of secondary cancer



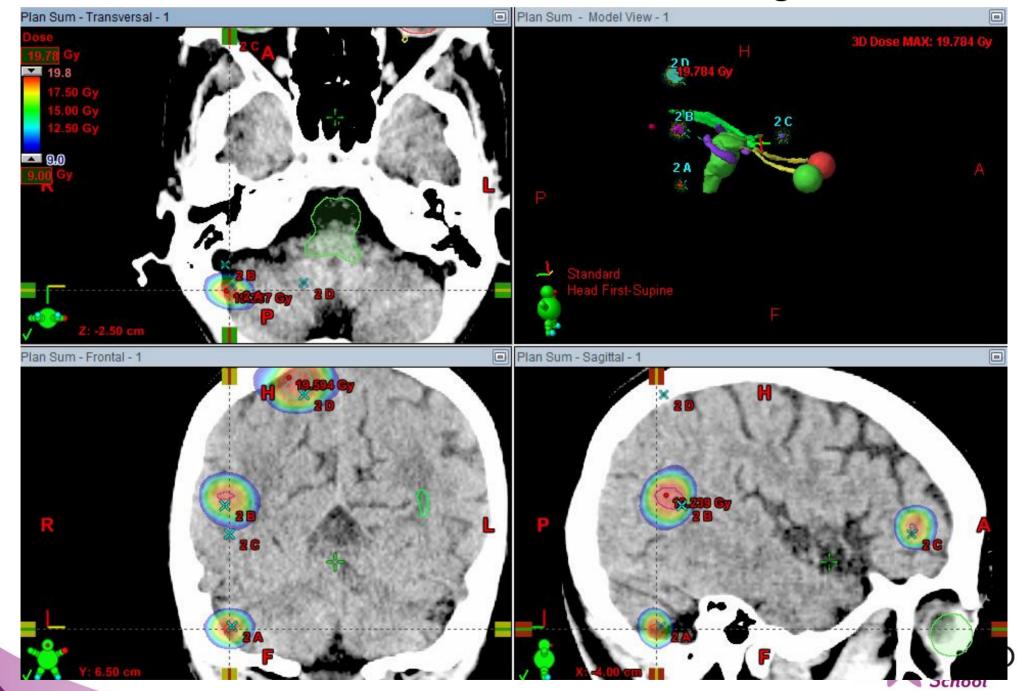
Stereotactic treatment – brain metastases



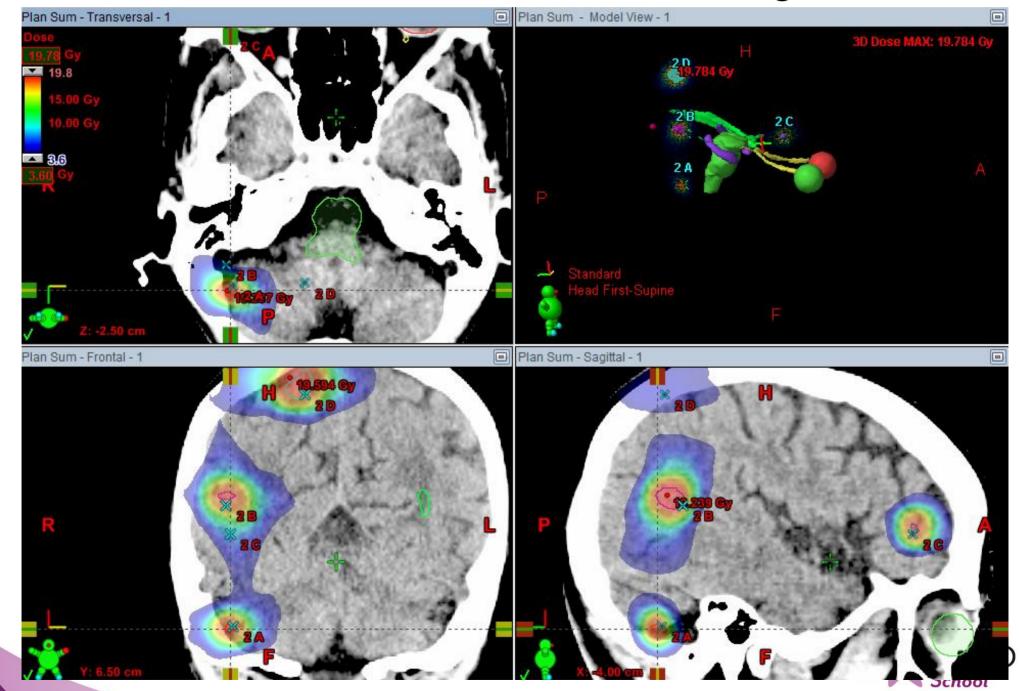
Stereotactic treatment – 4 targets!



Stereotactic treatment – 4 targets!



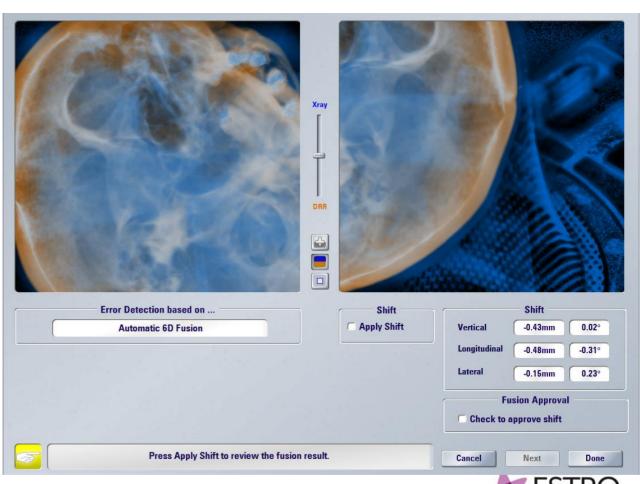
Stereotactic treatment – 4 targets!



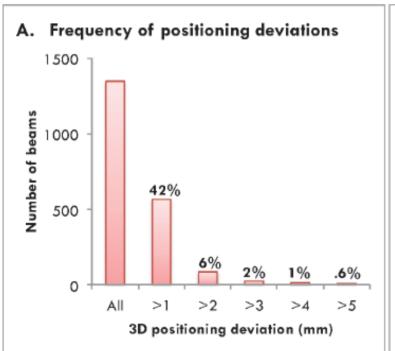
Delivery of stereotactic brain RT

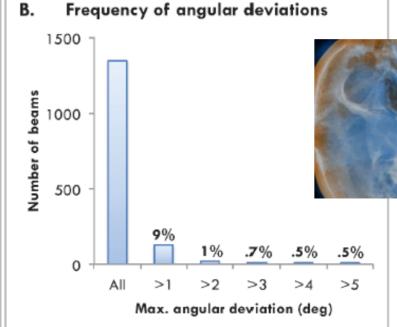
IGRT

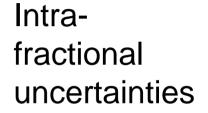
- Small PTV margins
- 6D corrections
- Rigs tolerance:
 - > <1mm
 - > <1°
- Non-coplanar RT delivery
 - Repeat imaging after couch rotation

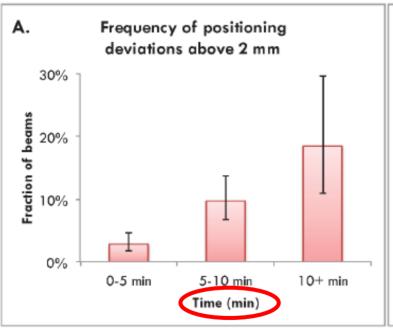


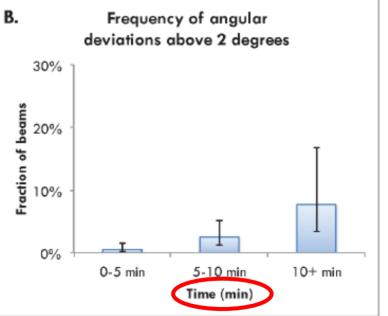








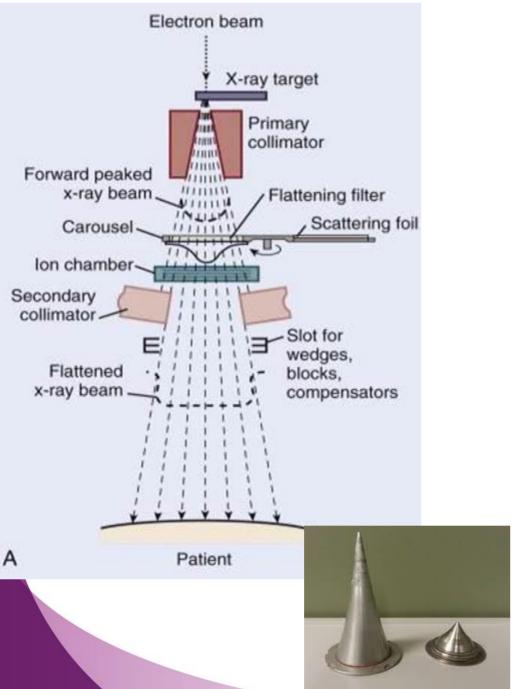




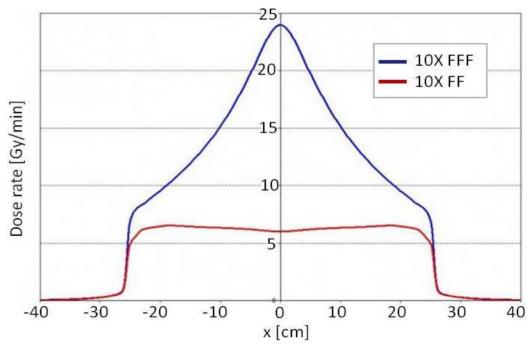


Tarnavski et al. Jour. of Radiosurgery and SBRT 2016

FFF – flattening filter free



Intensity modulated RT does not necessitate flat beams



FFF facilitates increase in dose rate & decrease in beam time by a factor of up to 6



A bit about the margins...

Margins depend on:

- RT technique
- IGRT strategy (Rianne's talk)

Example:

- 3DC RT & field verification at first treatment
 - > 5 mm CTV-PTV margin
- VMAT & daily IGRT with 6D:
 - ➤ 1-2 mm CTV-PTV margin



Considering the margins vs. daily IGRT workload



margins of 5 mm increase the treated volume by 50%





Brain

Rianne de Jong *RTT*, Academic Medical Centre, Amsterdam Prague 2017



Brain @the treatment machine

- How well can we set up the patient?
- How well can we image the target volume?
- How well can we correct the patient position?
- How stable is the patient position?
- Imaging dose for children

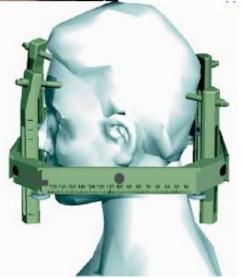


Commercial Immobilisation Options

- Thermoplastic mask
- Mask + bite block
- Frames
- Invasive frames













Set-up accuracy: interfraction motion ———— based on bony anatomy registration

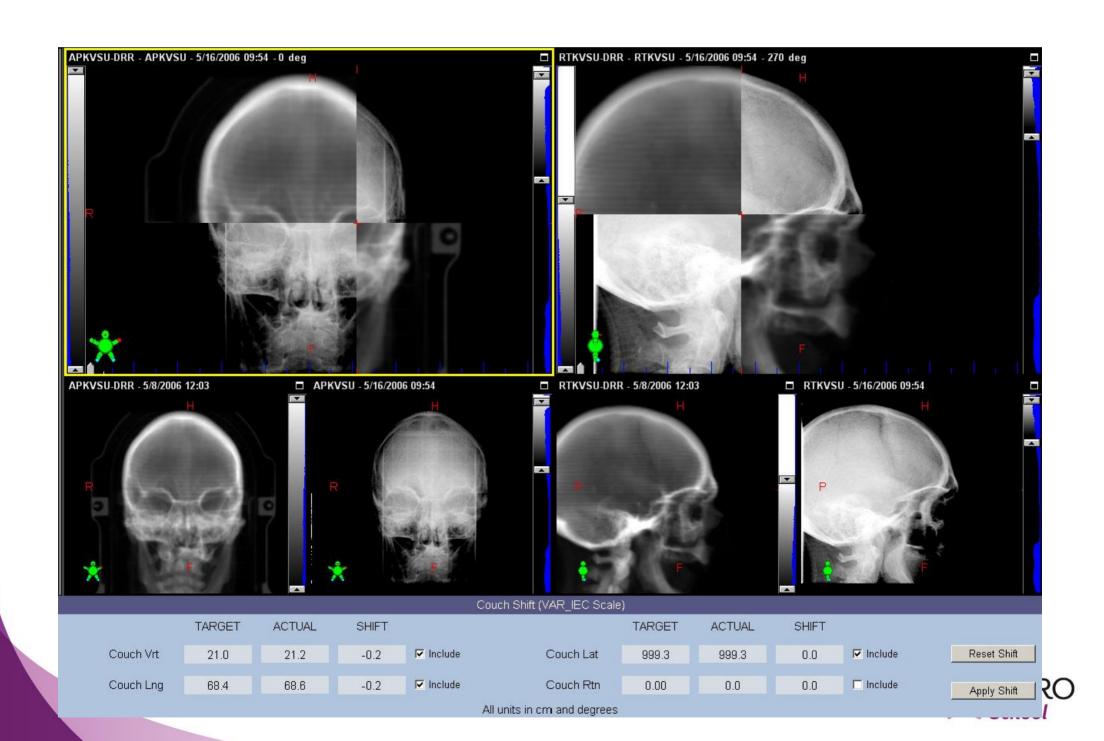
Study	Positioning system Imaging modality		Position error	
2D-2D image registration for verification of set-up				
Rosenthal 1995	Dental fixation	2.3mm ±1.6mm		
Sweeny 2001	Vogely Bale Hohner head Holder	Portal imaging	1.9mm ± 1.2mm	
Kumar 2005	Gill-Thomas-Cosman Portal imaging		1.8mm ± 0.8mm	
Georg 2006	Brain Lab Mask Portal imaging		1.3mm ± 0.9mm	
3D-3D image registration for verification of set-up				
Baumert 2005	Stereotactic mast	СТ	3.7mm ± 0.8mm	
Boda-Heggermann 2006	Scotch cast mask	СВСТ	3.1mm ± 1.5mm	
Guckenberger 2007	Scotch cast mask CBCT		3.0mm ± 1.7mm	
Masi 2008	Thermoplastic mask & Bite Masi 2008 block		2.9mm ± 1.3mm	
Bite-block		СВСТ	3.2mm ± 1.5mm	

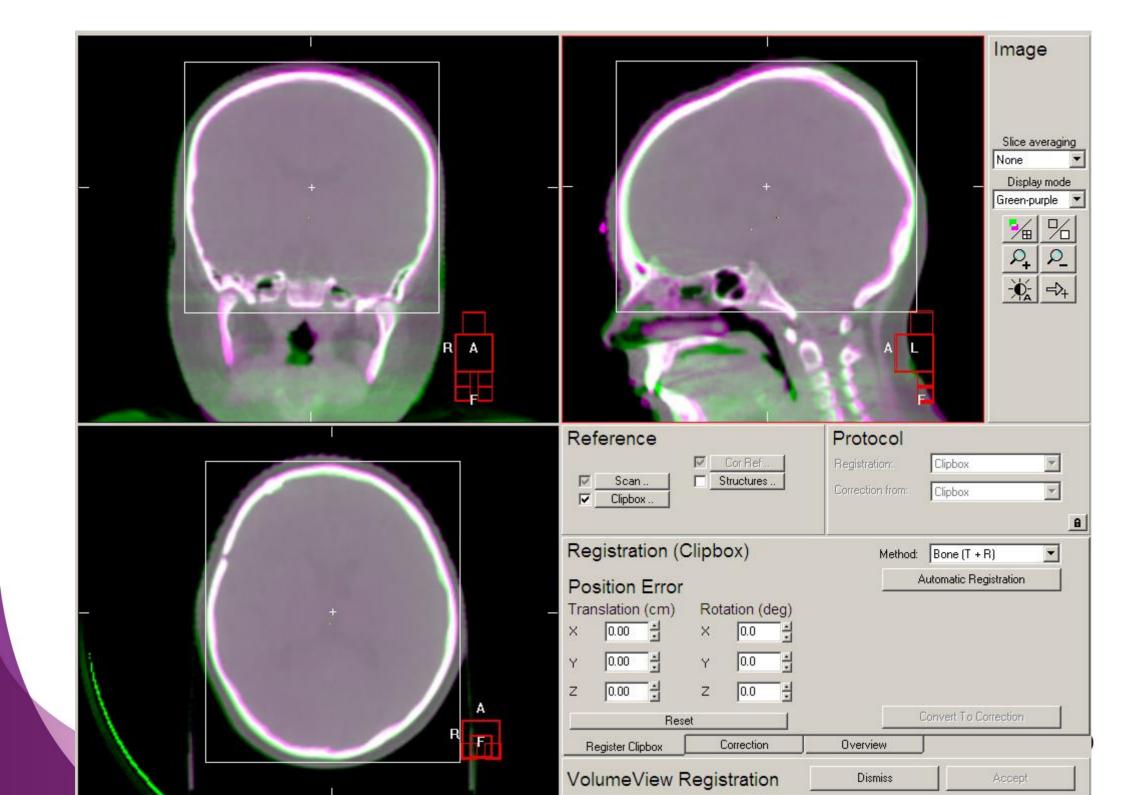


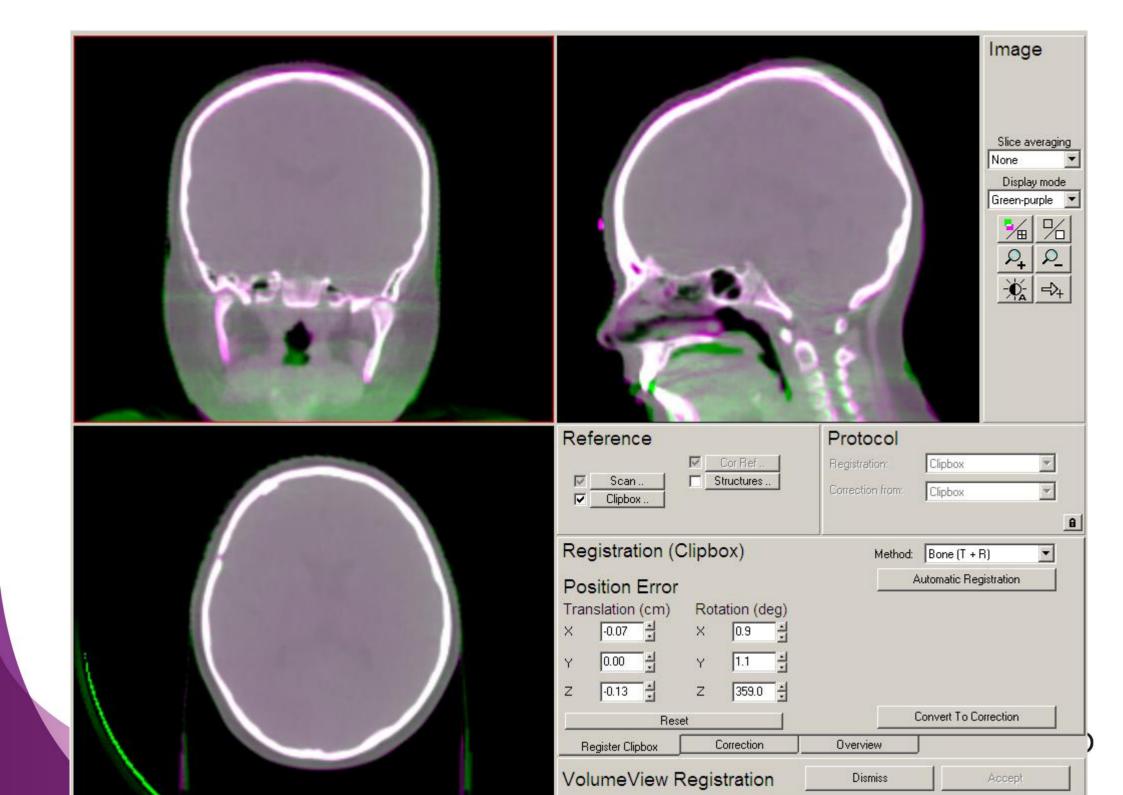
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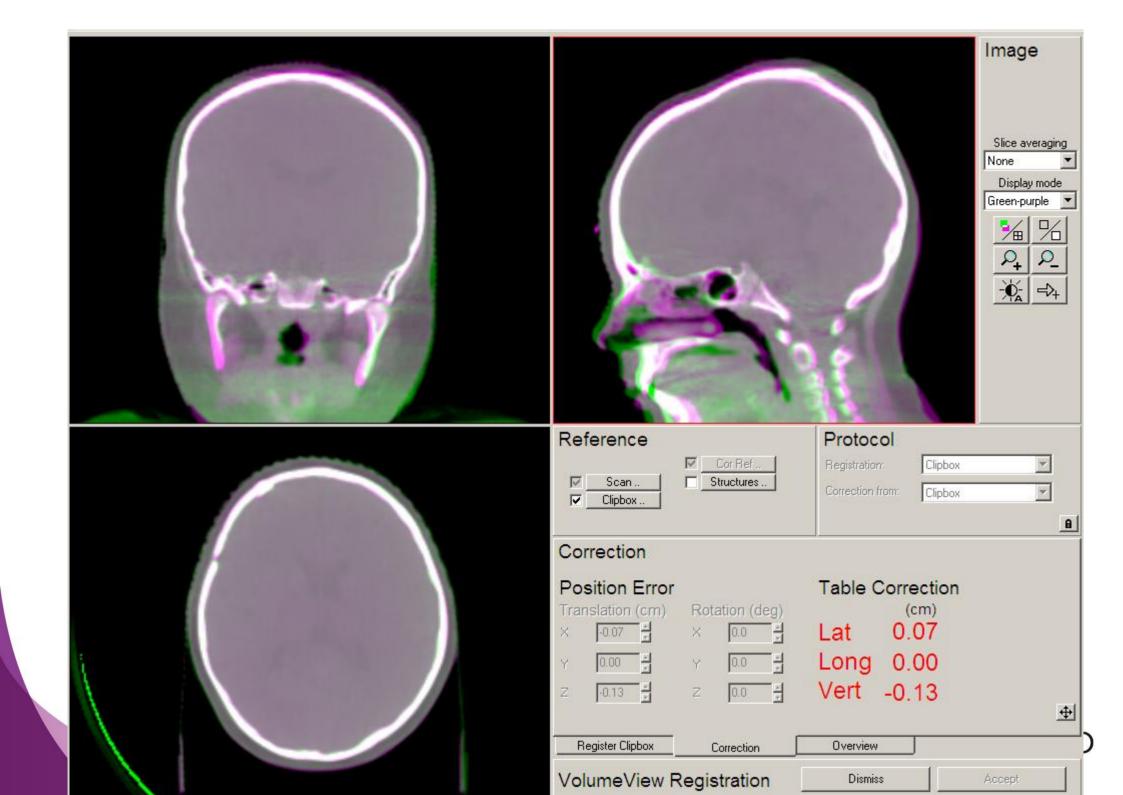
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	Bite-block	СВСТ	3.2mm ± 1.5mm	











Bony anatomy a good surrogate?

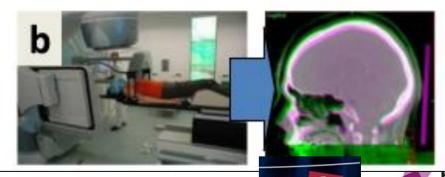
Internal motion of the intra cerebral tumor could be caused by

- Tumor progression
- Tumor shrinkage
- Changes of peritumoral edema

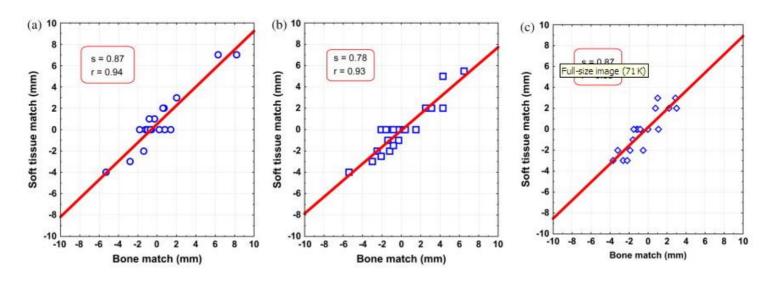
Set-up prior to treatment was verified based on the

- position of the metastasis (soft tissue match): imaging using an in-room CT scanner after application of iv contrast
- b) position of the bony anatomy (bone match): imaging using cone-beam CT





Bony anatomy a good surrogate?



	Difference between bone match and tumor match (mm)		
	LR	SI	AP
Mean ± SD	-0.5 ± 1.0	0.1 ± 1.1	-0.2 ± 1.0
Maximum	1.8	2.3	2



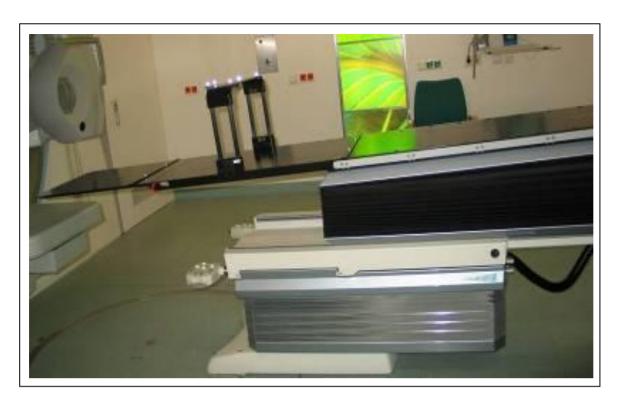
Bony anatomy a good surrogate?

Yes, but keep the time interval to a minimum!



------ How well can we correct errors?

Ask your physicist your table accuracy!



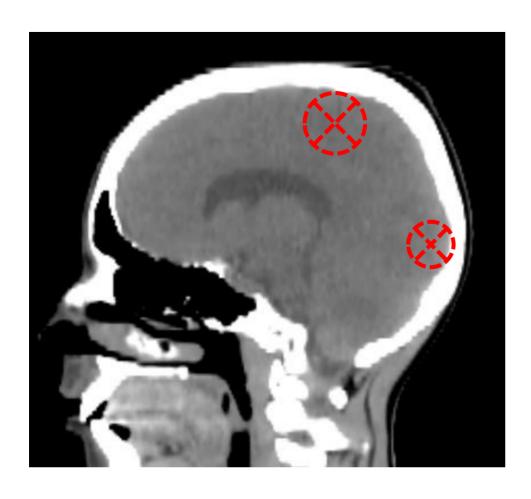
- Corrections up to
 3°
- Target is often spherical

Residual errors after image guidance with CBCT and robotic couch:

< **0.3mm** < **0.3**° Meyer 2008



Image registration How well can we correct errors?

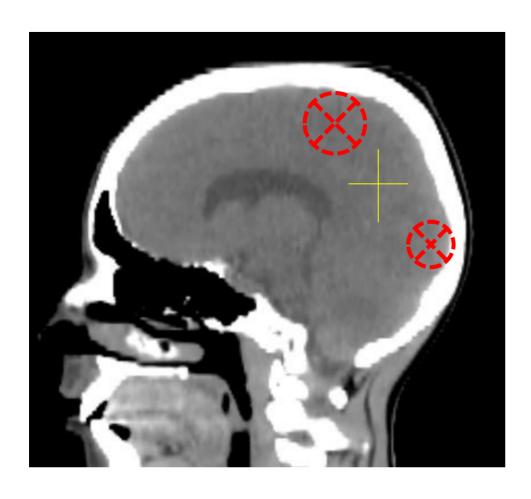


When multiple targets:

Rotations become important!



Image registration How well can we correct errors?



When multiple targets:

Rotations become important!

Use limit on rotations!

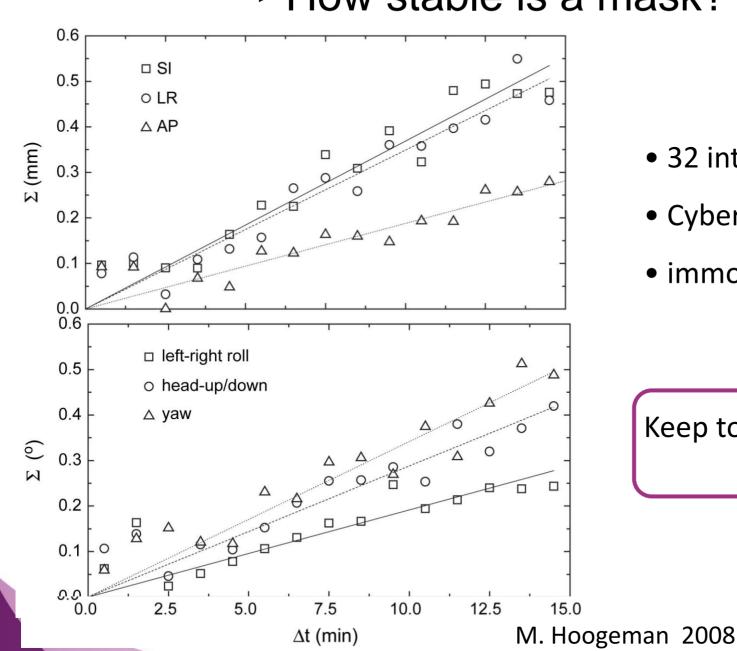


Correcting Patient position How stable is a mask?

Study	Immobilisation system	Imaging modality	Position error
Boda Heggeman 2006	Thermoplastic mask	СВСТ	1.8mm ± 0.7mm1.3mm
	Scotsch cast mask		1.3mm ± 1.4mm
Masi 2008	Thermoplasic mask & Bite block	CBCT	<1.0mm
	Bite block		<1.0mm
Lamda 2009	BrainLab mask	2D kV images	0.5mm ± 0.3mm
Ramakrishna 2010	BrainLab mask	2D kV images	0.7mm ± 0.5mm
Guckenberger 2007	Scotsch cast mask	CBCT	0.8mm ± 0.4mm
	Thermoplastic mask		0.8mm ± 0.5mm



Correcting Patient position How stable is a mask?



- 32 intracranial patients
- Cyberknife @ Rotterdam
- immobilized with a thermoplastic mask

Keep total treatment time as short as possible!



Margins for small leasions hypo fractionated

Adding up some/al the errors:

Delineation uncertainty 2 mm

Residual set up error after imaging (2D or 3D)

bone registration
 0.5 mm

soft tissue changes
 0.6 mm

Intrafraction motion 0.6 mm

PTV margin =
$$2.5 \Sigma + 0.7 \sigma$$

→ 1mm margin/0mm margin??

Literature show excellent local control!

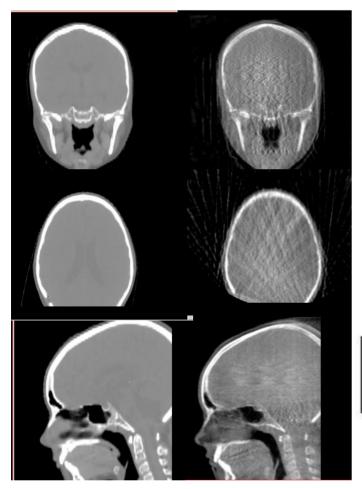


Imaging dose

——— Can we reduce dose for children?

100%, (0.5rpm)

8%



Lowest exposure settings

10ms & 10mA per projection

Using 'slice averaging' for display

	maximum deviation in outcome of registration			
	bony algorithm		grey value algorithm	
	translations (cm)	rotations (°)	translations (cm)	rotations (°)
skull	0.03	0.2	0.05	0.6
thoracic region	0.03	0.4	0.03	0.5
lumbar region	0.02	0.4	0.03	0.5



Imaging dose

——— Can we reduce dose for children?

Adult exposure 40ms, 32mA

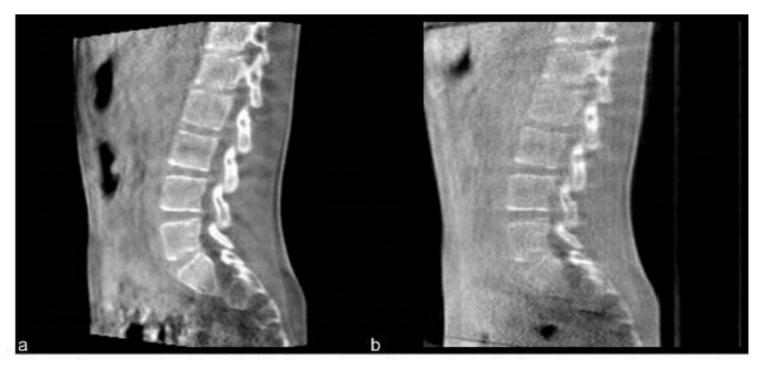


Kids exposure 10ms, 10mA



Imaging dose

——— Can we reduce dose for children?



40ms, 32mA, 0.5 rpm Display using slice averaging

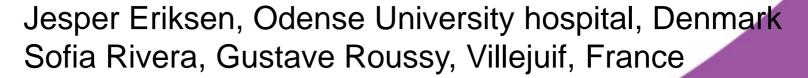
10ms, 10mA, 1.0 rpm Display using slice averaging





Case report: Head and Neck







Advanced skills in modern radiotherapy
June 2017

Changing traditional scenario in H&N cancer

- Increasing incidence of HPV positive tumors (+++ Oral Cavity)
- Improved outcome compared with HPV-negative tumors
- younger patients with limited comorbidity and good performance status, less likely to abuse tobacco and alcohol

Epidemiology of oral human papillomavirus infection

Christine H. Chung a,b, Ashley Bagheri a, Gypsyamber D'Souza c,*

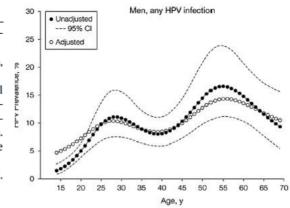
SUMMARY

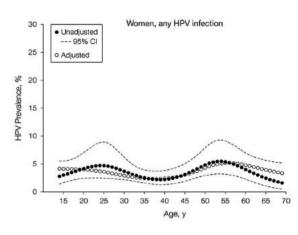
Objective: To describe what is known about the epidemiology of oral human papillomavirus (HPV) infection.

Methods: In this article we review current data on HPV prevalence, natural history, mode of acquisition, and risk factors for oral HPV infection.

Results & Conclusion: Over the past several years new studies have informed our understanding of oral HPV infection. These data suggest oral HPV prevalence is higher in men than women and support the sexual transmission of HPV to the mouth by oral sex. Data is emerging suggesting that most oral HPV infections usually clear within a year on and describing risk factors for prevalent and persistent infection. Recent data support likely efficacy of the HPV vaccine for oral HPV, suggesting vaccination may reduce risk of HPV-related oropharyngeal cancer.

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Patient history

Palate
Oral Cavity

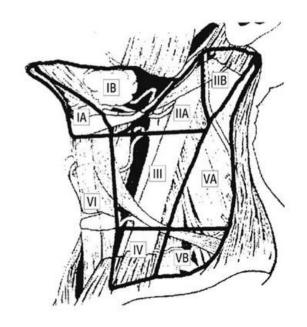
Pharynx
Epiglottis
Larynx opening into pharynx

Esophagus

- •60-year old man.
- 3 week history of nodal swelling, left side of the neck.
- No pain or dysphagia. No weight loss.
- No co-morbidity except from back pain.
- Ceased smoking in 1990, 10 pack-years.
- No daily use of alcohol.

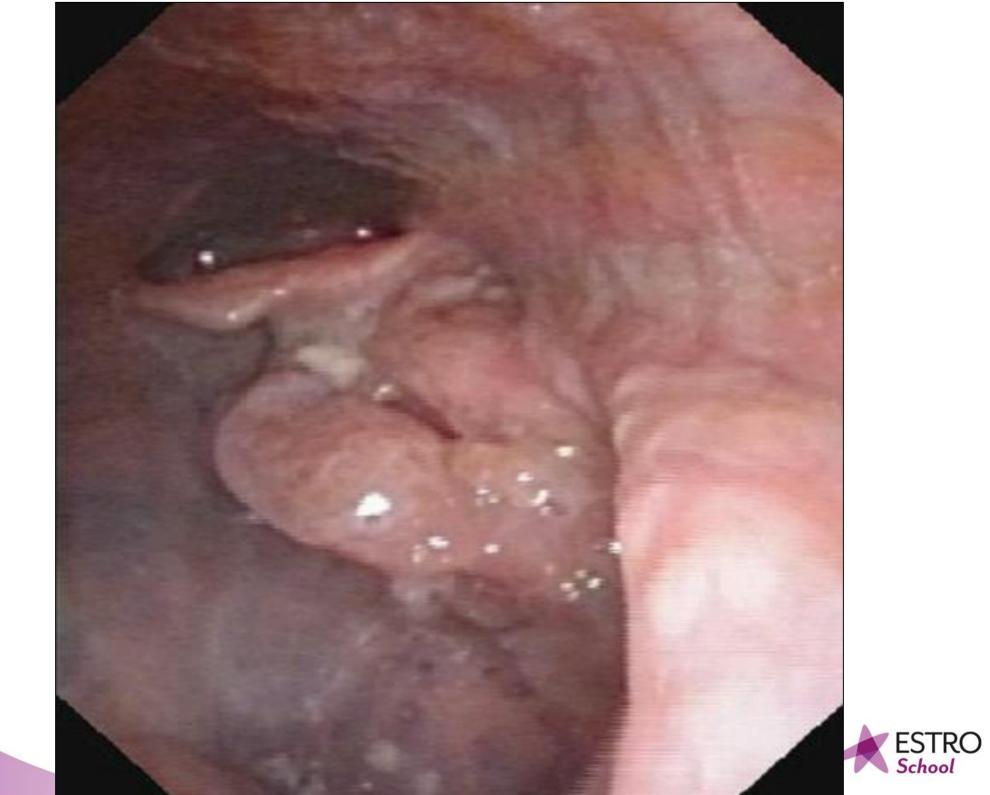


Clinical examination



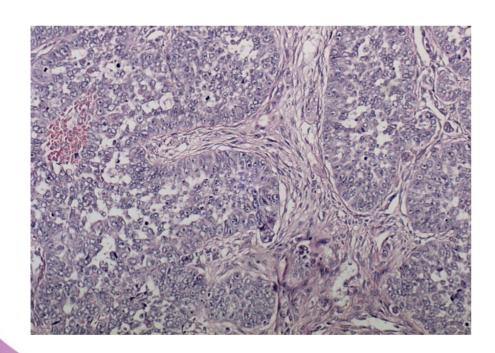
- Good performance (WHO PS 0)
- Base of tongue/vallecula area a 3x2x2cm large tumour is seen.
- •Proximal border of the tumour seems to be close to the lower pole of the left tonsil
- Otherwise normal fiber optic examination.
- Palpable node in region II, left side.
- Contralateral side normal.

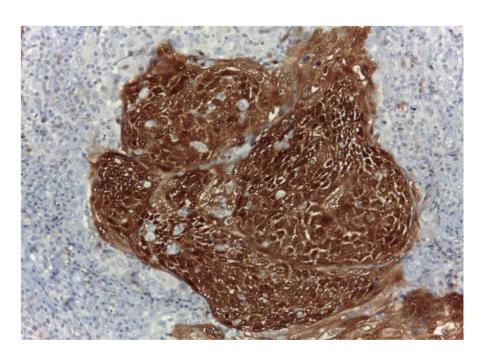




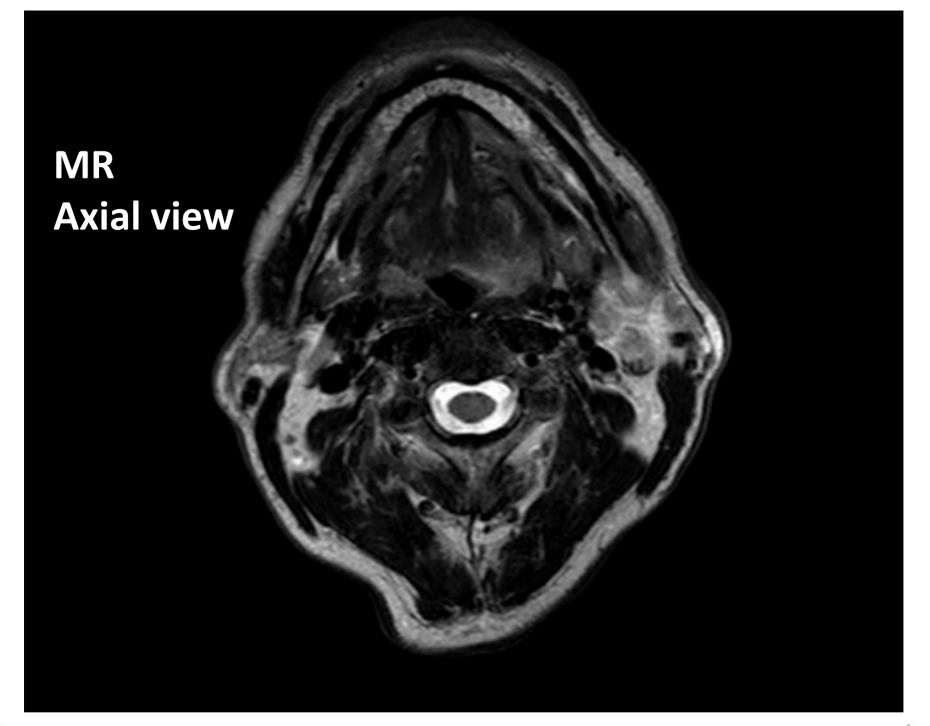
Pathology

- Moderate differentiated squamous cell carcinoma (G2).
- p16 positive (HPV marker)

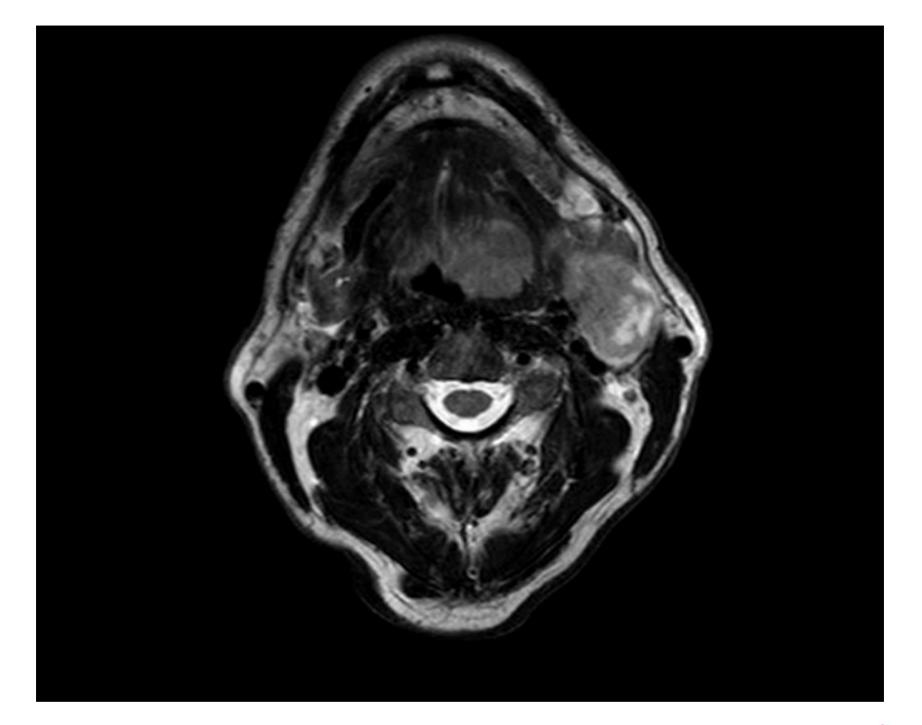




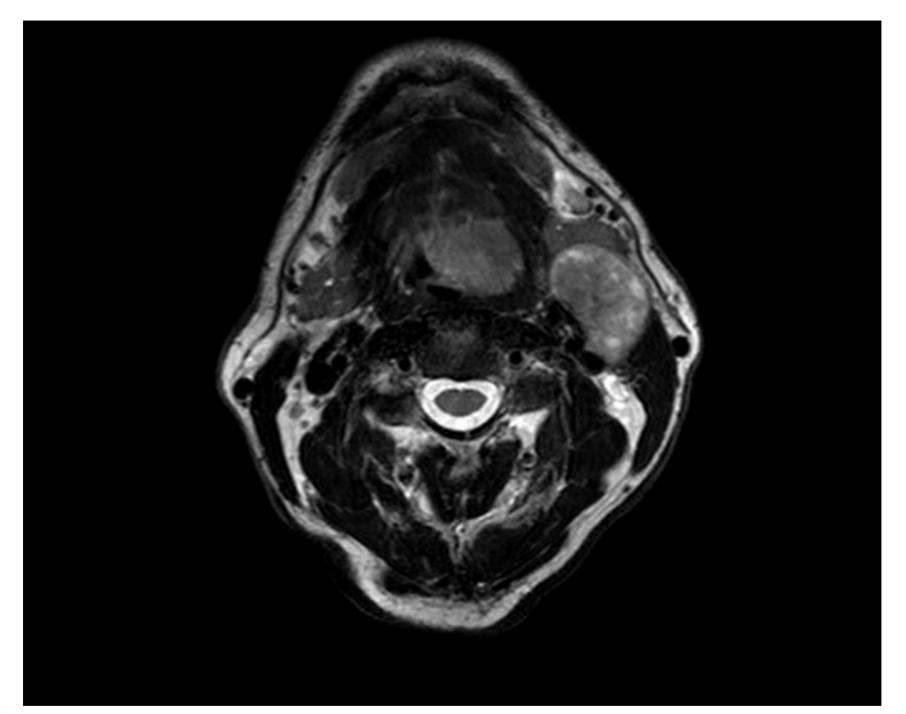




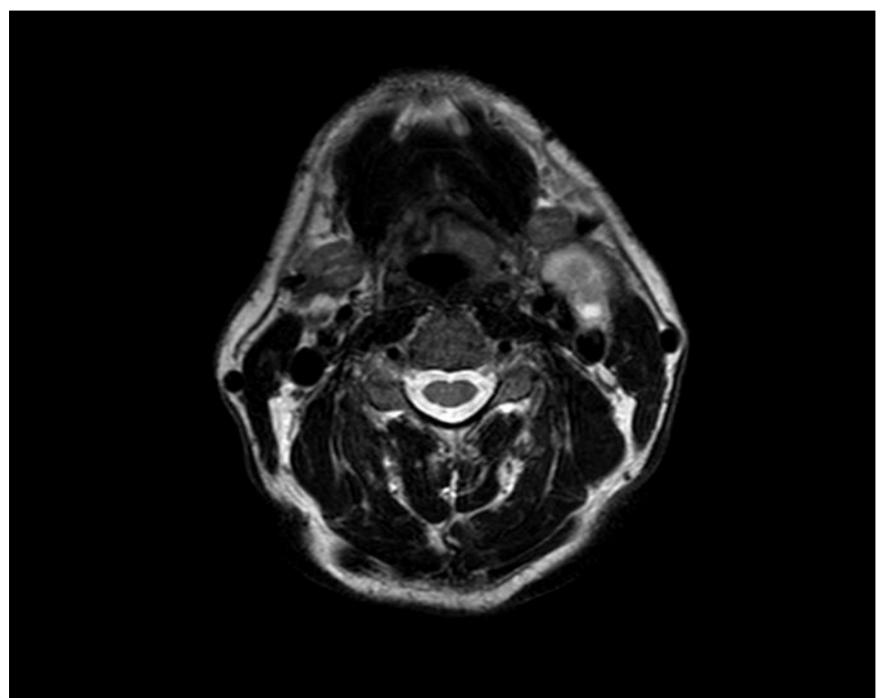




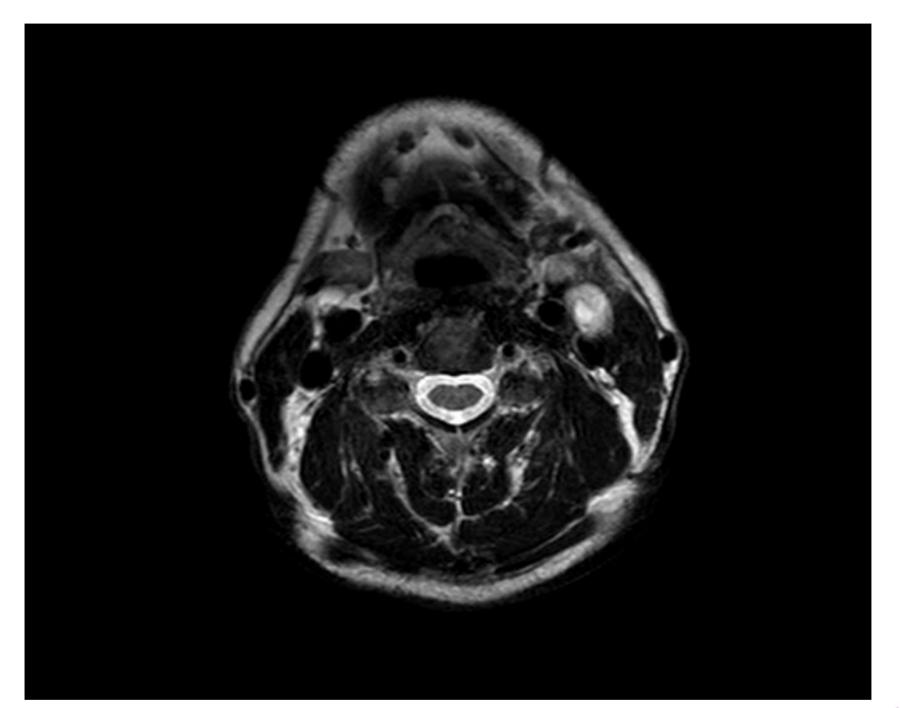




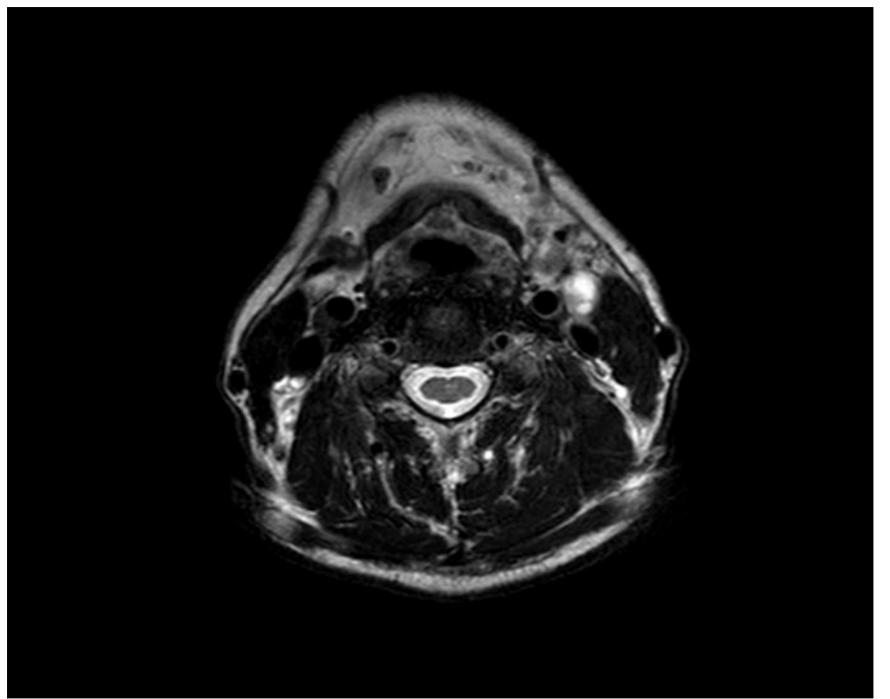




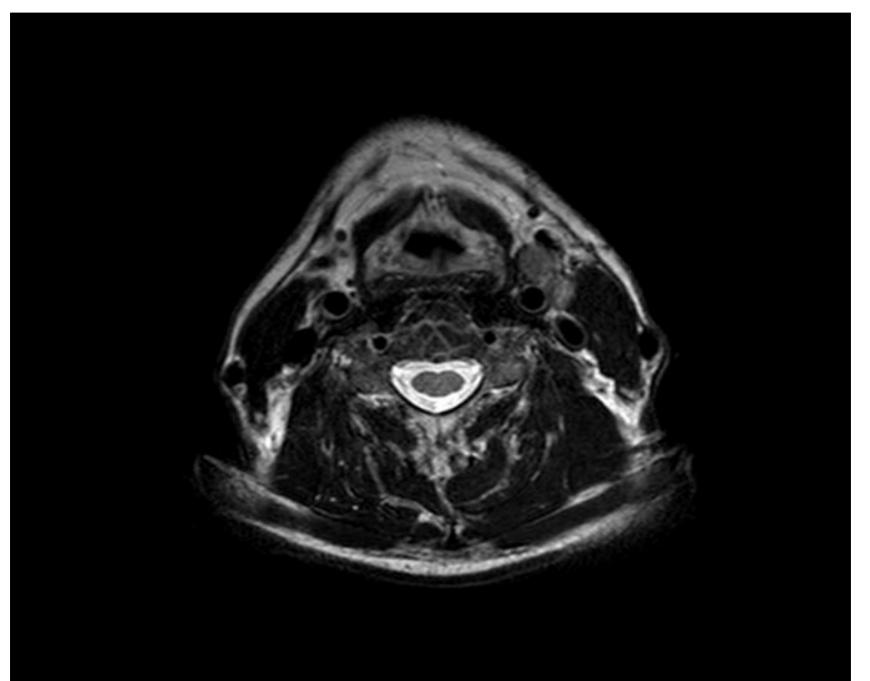




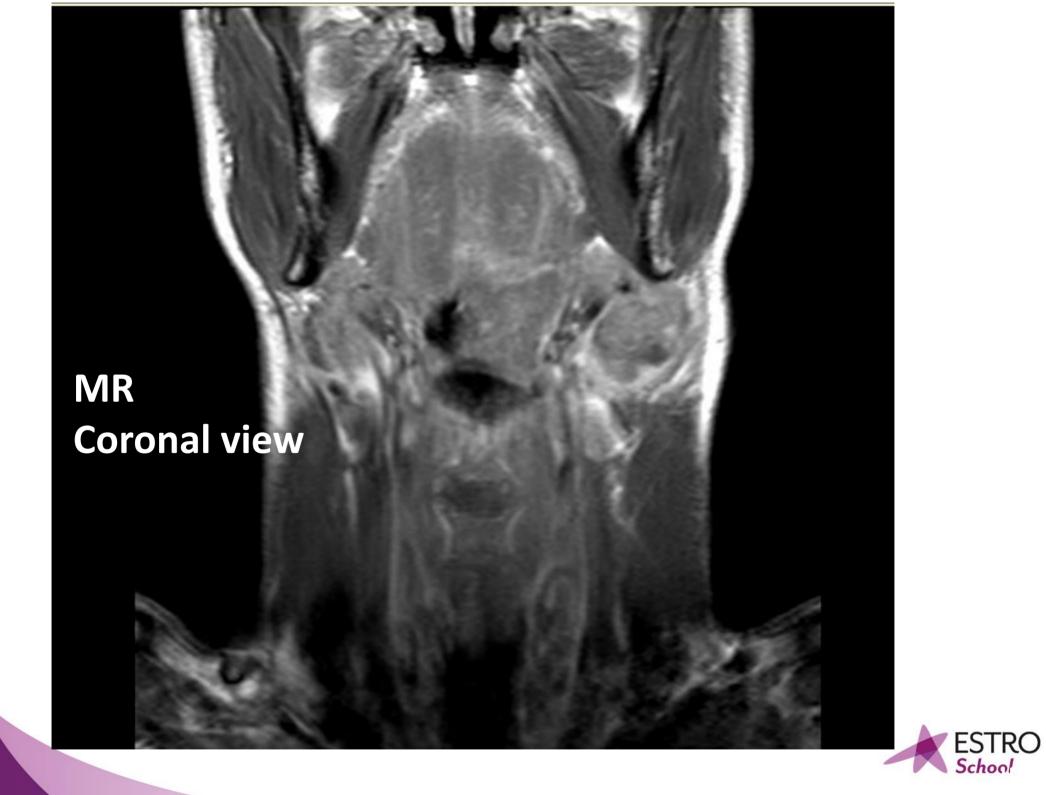


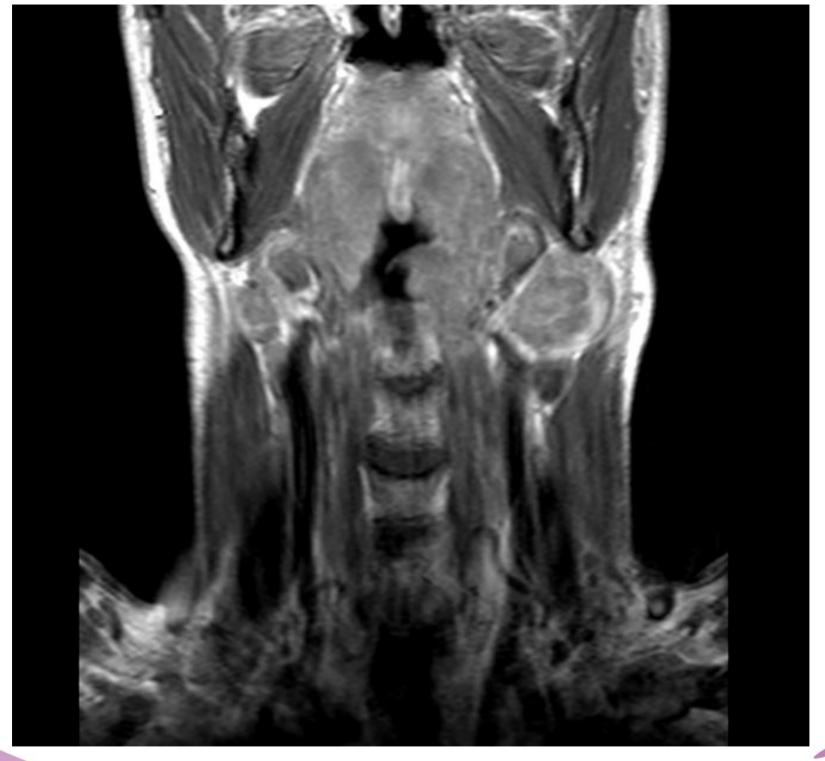








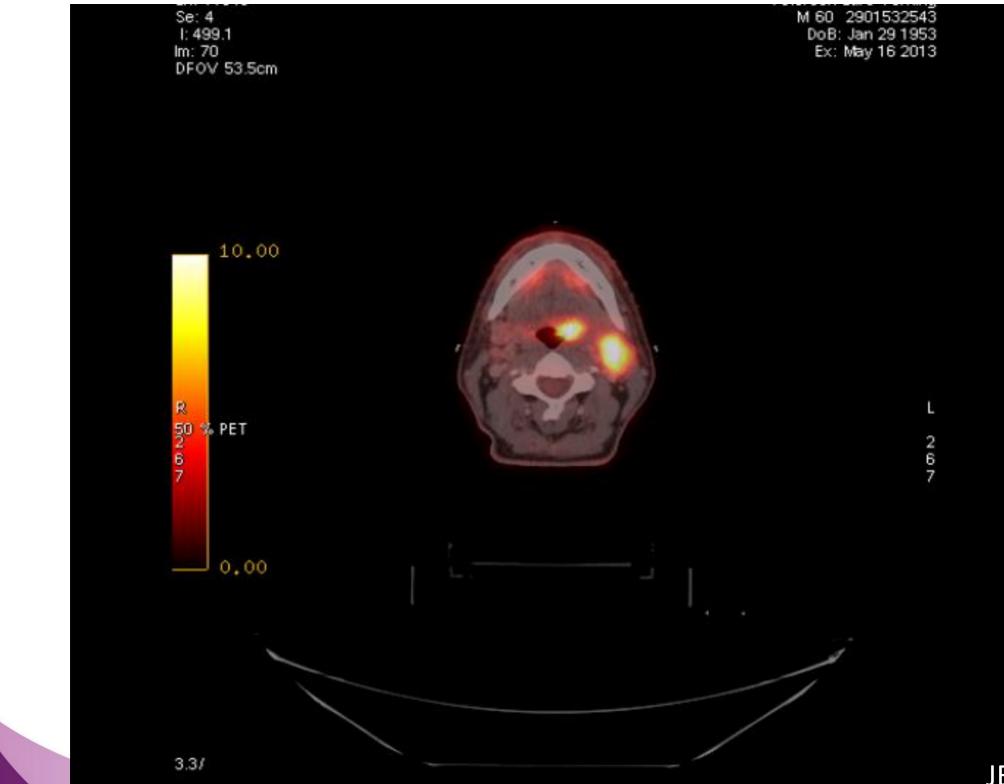


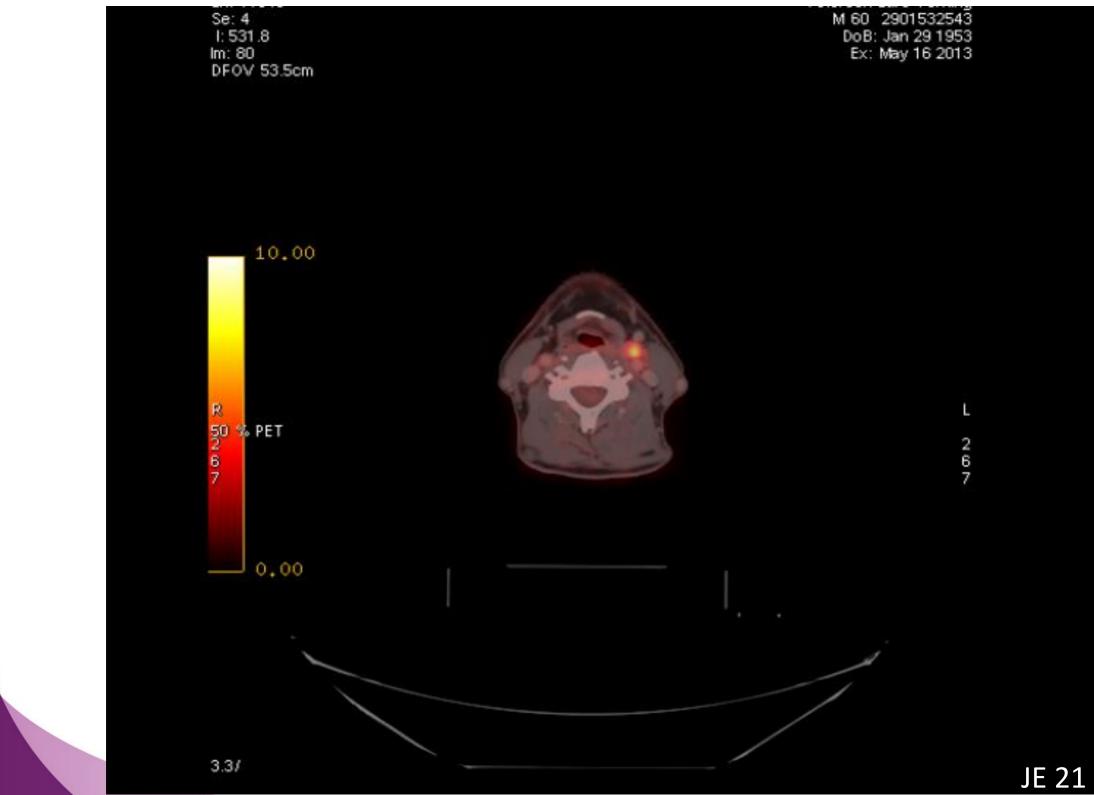


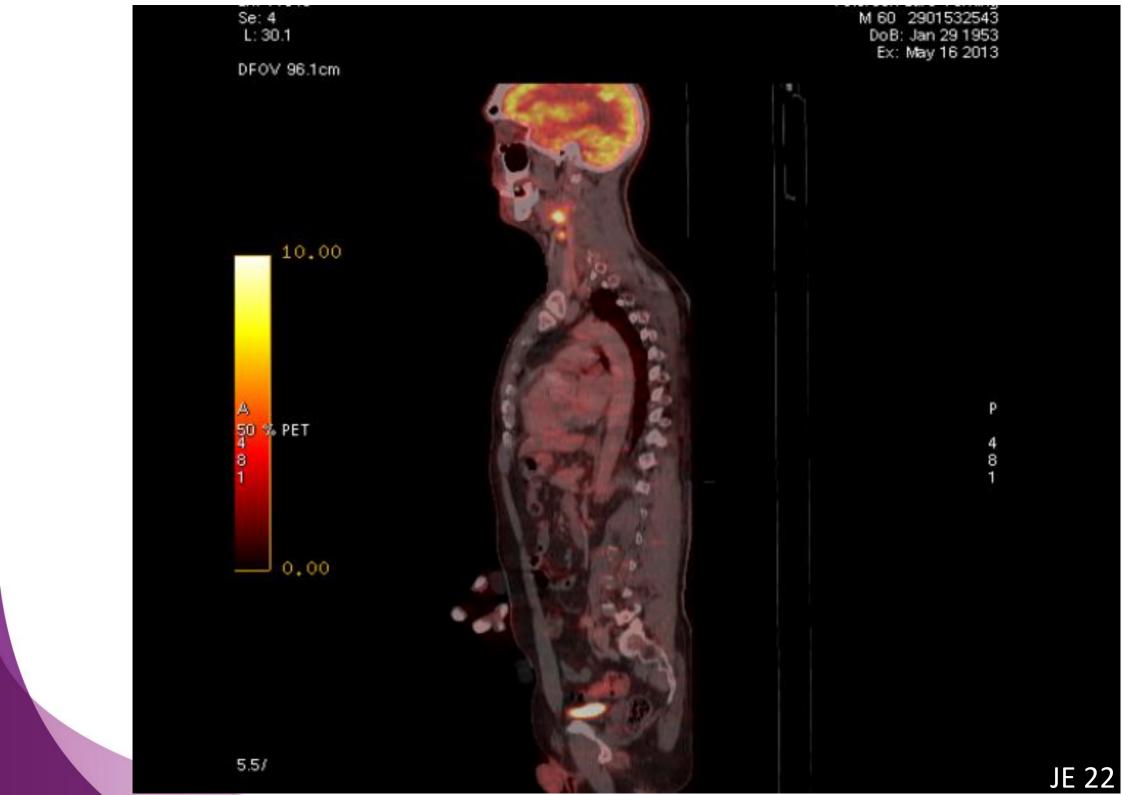






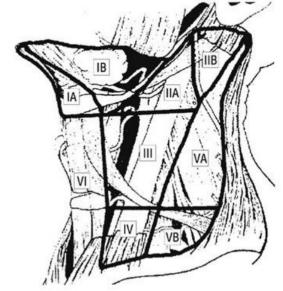






Ultrasound of neck

- One necrotic node in the upper part of left region II close to the submandibular gland; 3.5x2x2 cm.
- One node in left region III, 1.5x1x1 cm without preserved hilar region.
- Right side of the neck is normal.



Conclusions after diagnostic workup

- T2N2bM0 (stage IVa) SCC oropharyngeal tumour.
- Patient in a good performance with no relevant co-morbidity.



Treatment done

- 66 Gy/33 Fx; 2 Gy/Fx; 6 Fx/week.
- Concomitant weekly low-dose cisplatinum
 40 mg/m² (maximum 70 mg/m²).
- Concomitant hypoxic radiosensitization with nimorazole according to DAHANCA guidelines



Contouring guidelines

V. Grégoire et al. /Radiotherapy and Oncology xxx (2013) xxx-xxx

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Radiotherapy and Oncology xxx (2013) xxx-xxx

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Radiotherapy and Oncology

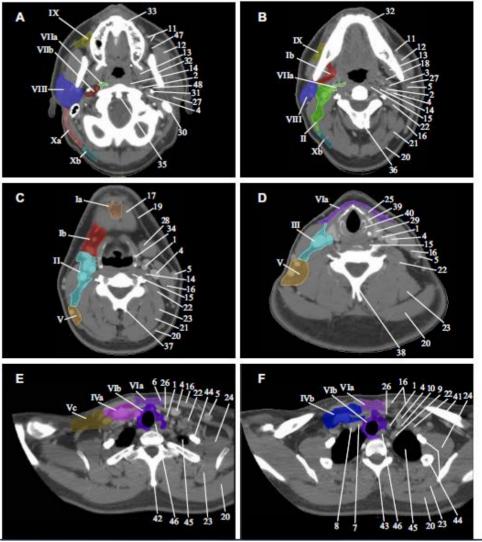
journal homepage: www.thegreenjournal.com



Original article

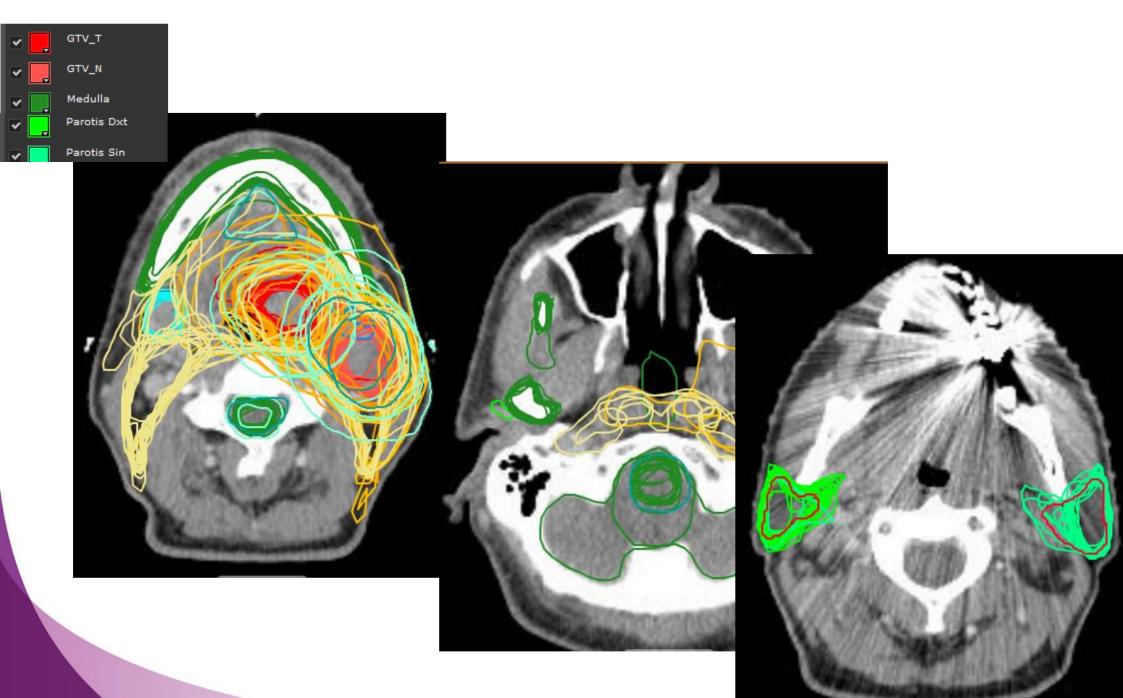
Delineation of the neck node levels for head and neck tumors: A 2013 update. DAHANCA, EORTC, HKNPCSG, NCIC CTG, NCRI, RTOG, TROG consensus guidelines $^{\dot{\approx}}$

Vincent Grégoire ^{a,*}, Kian Ang ^b, Wilfried Budach ^c, Cai Grau ^d, Marc Hamoir ^e, Johannes A. Langendijk ^f, Anne Lee ^g, Quynh-Thu Le ^{h,i}, Philippe Maingon ^j, Chris Nutting ^k, Brian O'Sullivan ^l, Sandro V. Porceddu ^m, Benoit Lengele ⁿ





Case used for H&N Falcon online WS



Take home messages:

- HPV positive tumors are changing H&N cancer traditional scenario
- Positioning remain key points for these highly conformal treatments (IMRT+++)
- Target and OAR contouring remains an issue: Highly heterogeneous contours
- Crucial need for contouring guidelines and training





H&N case - Physics considerations

Peter Remeijer



Head and neck

 Complex target volume (GTV, CTV, lymph nodes, PTV)

- Many organs at risk (OARs)
 - > Spine
 - Brainstem
 - Salivary glands
 - **>** ...
- OARs often close to PTV



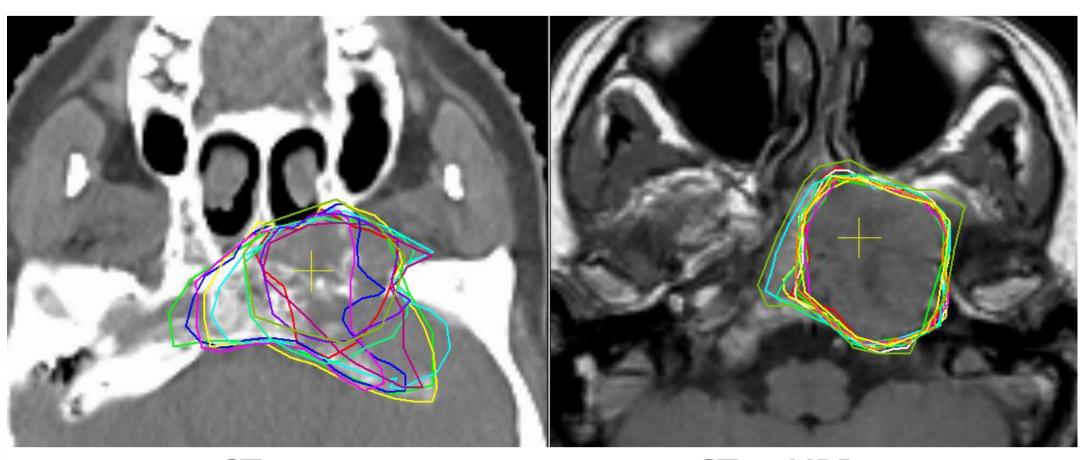
Delineation and planning

- Multi-modality imaging (PET/MR) has a large impact on delineation uncertainties
- Image registration becomes important
- PET/MR scans preferably in treatment pose
 - Minimizes errors due to deformations





Multiple modalities improve H&N delineation



CT

SD 4.4 mm

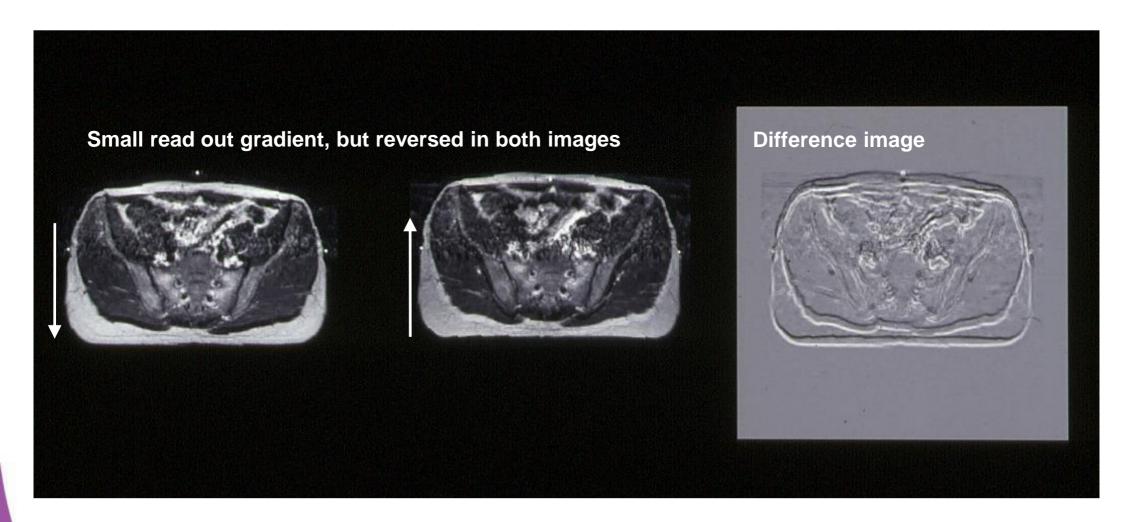
CT + MRI

SD 3.3 mm



Steenbakkers et al

MRI artifacts can cause invisible geometrical errors!



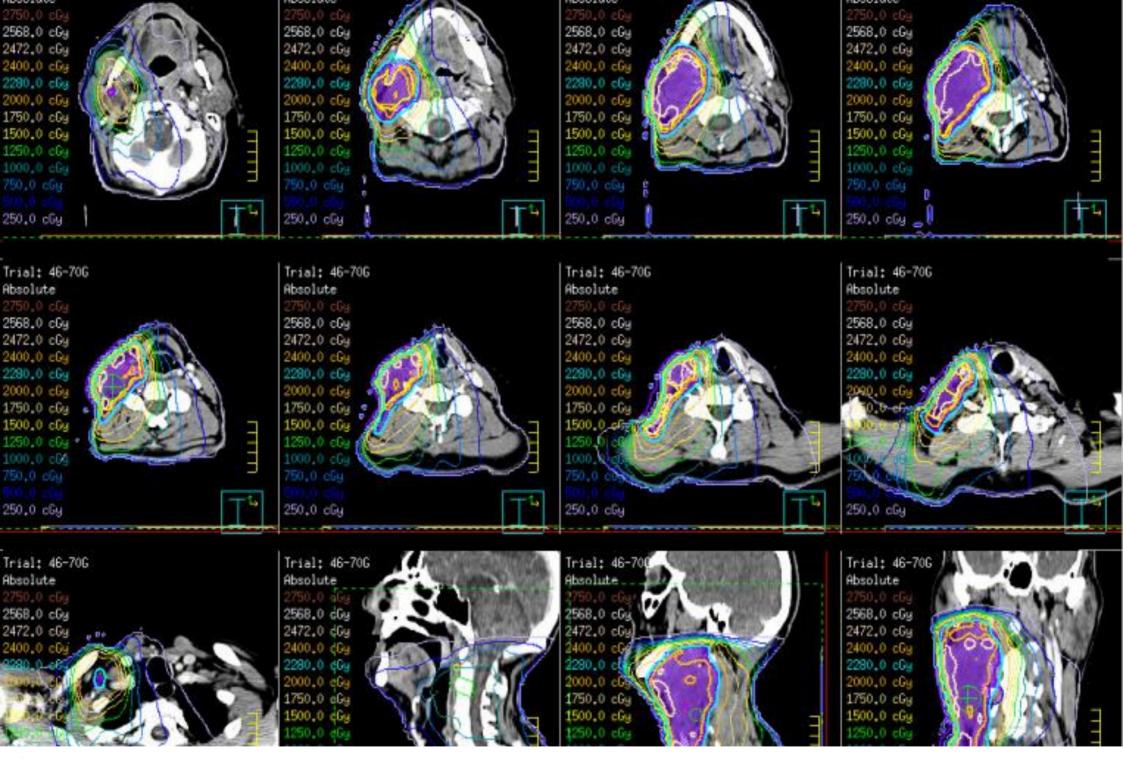
What you see is not always what you get



Delineation and planning

- Multi-modality imaging (PET/MR) has a large impact on delineation uncertainties
- Image registration becomes important
- PET/MR scans preferably in treatment pose
 - Minimizes errors due to deformations
 - Focus on the main region of interest
- Planning: IMRT or VMAT





Complex planning techniques \rightarrow Margin becomes more critical

Treatment - All is well?

# Patients	# Images	Direction	σ _{sys} (mm)	σ _{random} (mm)	Reference
31		ml	1.8	1.5	(Bel et al., 1995)
		СС	1.7	1.1	
		ap	2.0	1.6	
26	356	ml	1.8	1.6	(Vos et al., 1997)
		СС	2.7	1.5	
		ap	1.7	1.2	
		rot ant	1.2°	0.8°	
		rot lat	0.7°	1.0°	
12	192	СС	2.0	1.4	(Yan et al., 1997b)
		ap	1.3	1.7	
12	290	ml	1.8 ^a	1.4	(Gildersleve et al., 1995)
		СС	2.2 ^a	1.4	
		ap	1.7 ^a	1.4	

Portal imaging data from a long time ago...

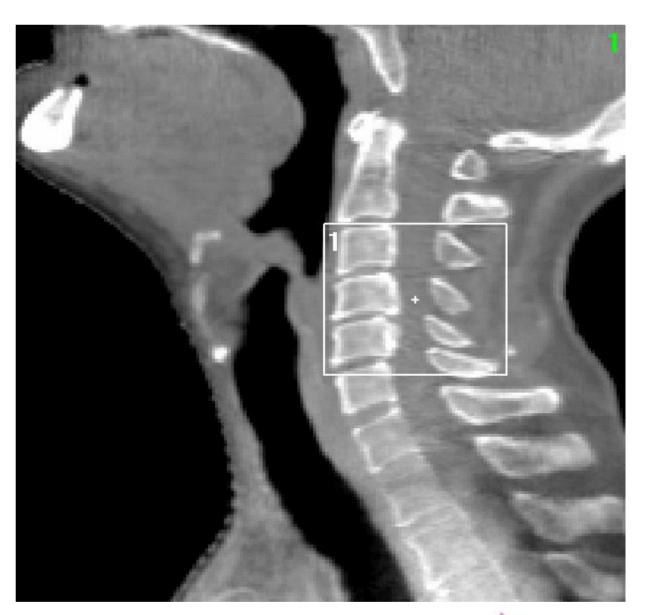
- Systematic errors: 1 to 2 mm (SD)
- Random errors: 1 to 2 mm (SD)



Cone beam CT → Shape changes!

- Pose
- Weight loss
- Tumor regression
- •

→ Non rigid





Treatment – image registration

Single ROI

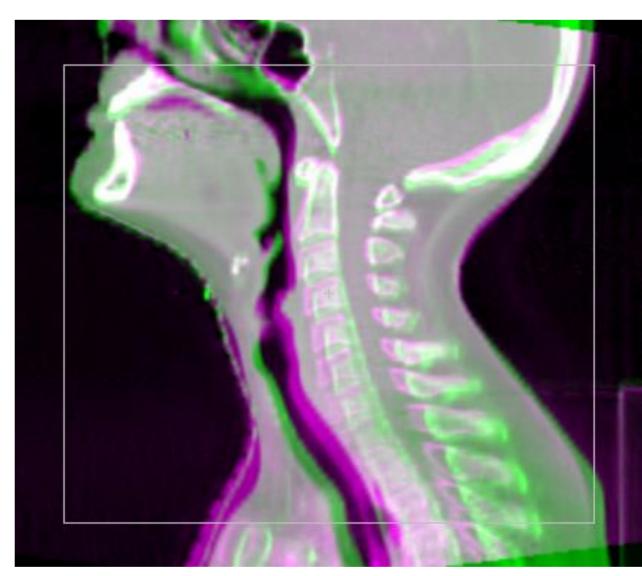
ROI encompasses:

- PTV
- Vertebrae
- Base of skull
- Jaw

Purple: CBCT

Green: planning CT

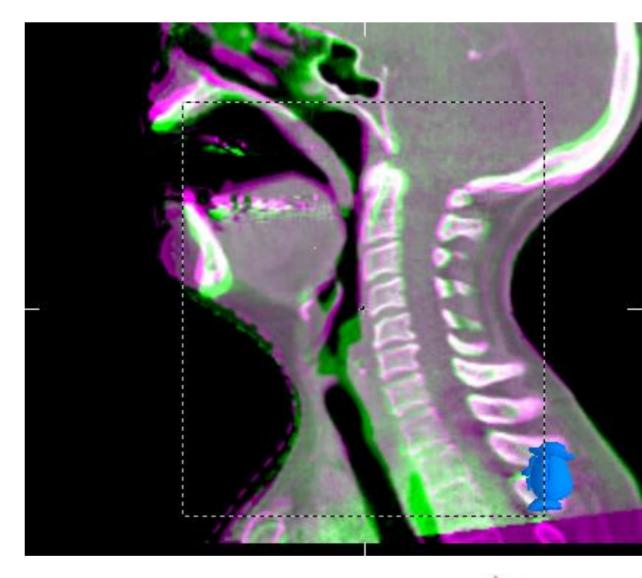
Overlay: white = match





Single ROI registration

- Match inaccurate
 - Misregistration?
 - Deformation?





Use multiple ROIs

Allows:

- Accurate local registration
- Assessment of local setup errors

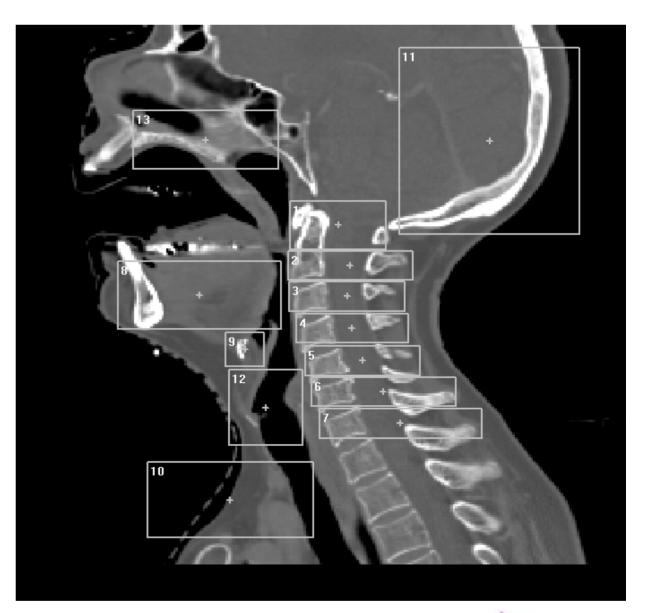
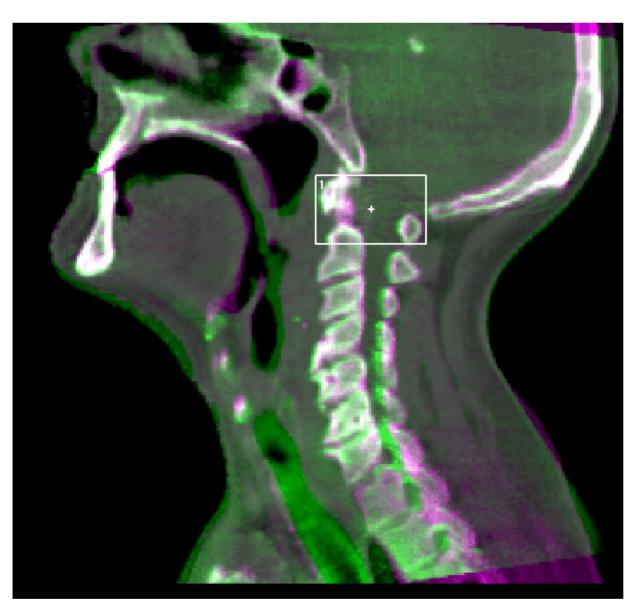




Image registration



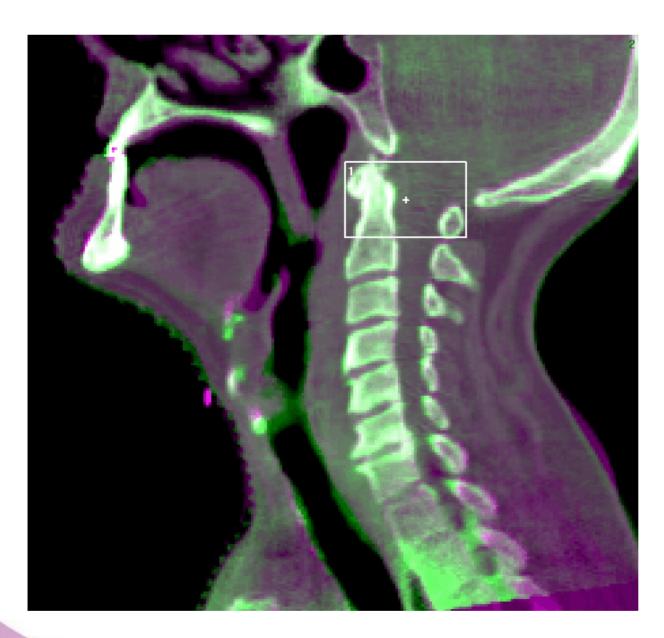
- bony anatomy registration
- Loop over ROIs

Purple: planning CT

Green: CBCT



Validation of registration

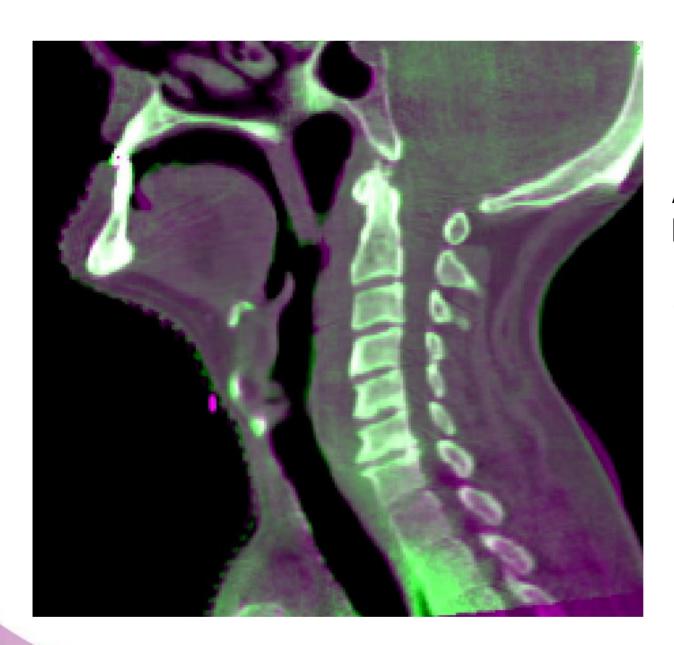


All registrations separately

→ Easy



Validation of registration



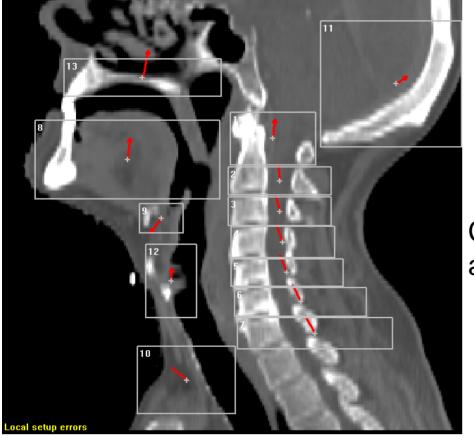
All registrations at once by warping

→ Fast



Corrections

Couch shift: Average setup errors

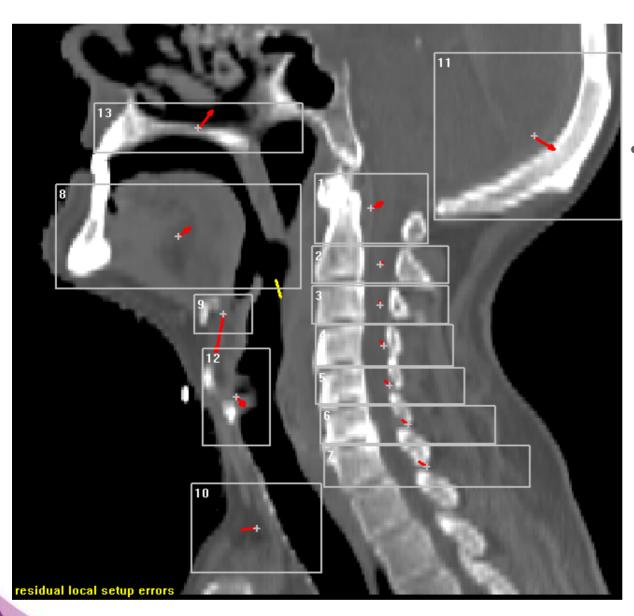


Correct average





Corrections



Residual errors!



Corrections

Residual errors

Warnings:5 mm or 5°

Overview				Residuals after correction			
Clipbox 2	-0.31	0.31	0.04	5.1	-4.4	-0.1	
Clipbox 3	-0.25	0.20	0.07	3.3	-3.7	0.0	
Clipbox 4	-0.10	0.05	0.21	0.6	-2.1	0.9	
Clipbox 5	0.11	-0.13	0.54	-2.6	-0.2	1.9	
Clipbox 6	0.14	-0.17	0.61	-3.0	0.5	1.6	
Clipbox 7	0.23	-0.14	0.59	-2.8	0.5	2.3	
Clipbox 8	0.04	0.02	-0.18	1.4	1.5	0.1	
Clipbox 9	0.02	0.16	8.02	5.5	0.4	0.3	
Cliphou 10	In no	0.11	0.00	0.5	1.4	0.1	

- 3 consecutive warnings:
- → Evaluate → Possible re-plan



Alternatives?

- If only one region of interest
 - ➤ Limit size to most important structures e.g. the boost area
 - > If deviations are visible outside this area...
 - → Retrospectively register and discuss



- If two regions of interest (Dual registration in XVI)
 - ➤ Use one region of interest for most important structures e.g. the boost area
 - Use other region of interest for larger area and specify tolerance limits. When limit is exceeded...
 - → Discuss
- Re-plan if deformation is persistent \rightarrow ART



Margins

- Delineation: 2-3 mm SD
- Setup: 1-2 mm SD (Portal imaging)

	Systematic	Random		
Delineation	2-3 mm	_		
Setup	1-2 / 1-4 mm	1-2 / 1-4 mm		
Organ motion	Depends on tumor location	Depends on tumor location		
Total	2-4	1-4 mm		

- Margin: 5 mm (best case) 13 mm (worst case)
- E.g. boost area → high precision/small margins
 Nodal regions → less precision/but CTV!



Take home messages

- Head and neck is a complex site
- Geometrical uncertainties are underestimated
 - Especially with portal imaging
 - Deformations are an important factor
- Margins will depend on correction strategy
- For persistent anatomical changes replanning is a good option
- Search for best correction strategy ongoing ©





Head and Neck IGRT: An RTT Perspective



Liz Forde, RTT
Assistant Professor
The Discipline of Radiation Therapy
School of Medicine
Trinity College Dublin







Fundamental IGRT Questions

- When should I image?
 - Frequency
- **How** should I image?
 - > Technology
 - > Projection
- What can I see?
 - What is my target
- What should I match to?
 - Surrogate for target position





Site Specific Points to Consider

- The head and neck is a regions rich in radiosensitive structures (serial organs)
- Margins are typically tight
 - > 0.3cm -0.5cm
- IMRT or VMAT are now standard and carry with them highly conformal dose distributions and multiple targets



Site Specific Points to Consider

• In addition to standard match structures also review:

- Position of mouth bung (if used) is correctly in place
- Bolus is positioned correctly (no gaps)
- Change in tumour size



Site Specific Points to Consider

- Gaps between skin and mask
- Shoulder position
 - Neubauer *et al.*, 2012
- Direct clinical impact of translations and rotations have on adjacent structures
 - > True OAR
 - OAR PRV



Pre Treatment

CT Simulation

Slice thickness

- Accurate delineation
- Accurate dose calculation
- Improved DRR resolution
- 2.5-3.0mm

Registration of diagnostic imaging

Contrast

IV

No pre contrast scan

Bolus

Scan with bolus on

Planning

3DCRT
IMRT Standard for this patient group

Beware the steep dose gradients

Shoulders Avoid?



Match Anatomy

- Bony landmarks
- Vertebrae
- Angle of mandible
- Orbital rim
- Frontal sinus
- Pituitary fossa



2D

- MV (EPI) is adequate for visualisation of bony anatomy
- Single projection not recommended for H&N
- Need to confirm isocentre in two planes
- Of less value when treating with IMRT
 - Field borders
 - Ciao images
- Impact of dose when imaging daily with MV

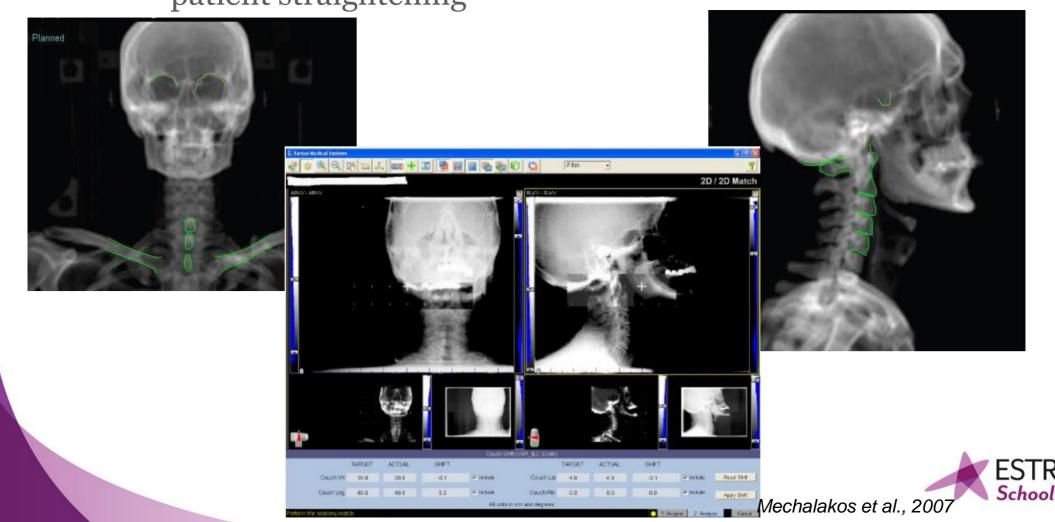




2D/2D

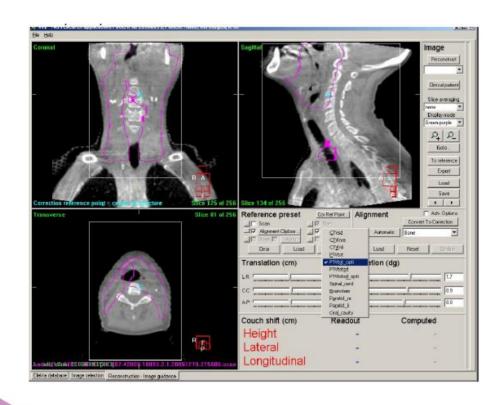
Improved visualisation and image quality

 Large FOV assess anatomy across whole target volumes and patient straightening



3D

- Peter has covered this in excellent detail!
- Consider other structures to review
 - > 45Gy isodose line



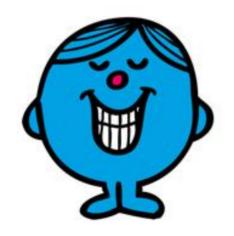




All Very Straightforward!

• But wait...there's more...

MR. PERFECT



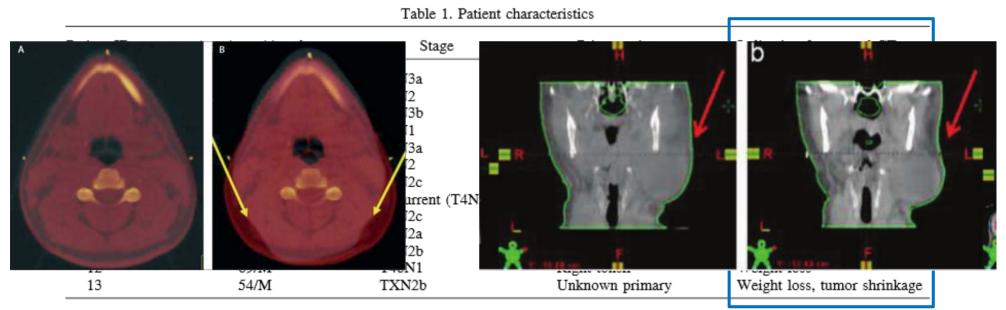
MR.WRONG



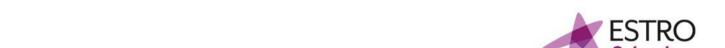


Tumour Shrinkage and Weight Loss

- Despite nutritional support these patients typically suffer significant weight loss during treatment
 - Impact on setup accuracy
 - Role of prophylactic PEG



Abbreviations: ID = identification number; M = male; F = female; NPX = nasopharynx; BOT = base of tongue. * Patient died of pneumonia after completing 23 fractions.



Tumour Shrinkage and Weight Loss

Replanning during IMRT for H&N cancer • E. K. HANSEN et al.

Dosimetric Impact!

Table 3. Dosimetric comparisons of the 2nd portion of treatment with and without replanning

	-		portion of treatment	
Dosimetric end point (mean values)	(1st CT/1st plan)	Replanned (2nd CT/2nd plan)	Not replanned (2nd CT/1st plan)	p value
PTV_{GTV}		Γ		
D ₉₉	38.1 Gy	28.3 Gy	26.0 Gy	0.05
D_{95}^{95}	40.3 Gy	30.3 Gy	28.1 Gy	0.02
V_{93}^{93}	99.5%	99.4%	92.5%	< 0.001
PTV _{CTV}				
D_{99}	30.9 Gy	22.9 Gy	18.3 Gy	< 0.001
D_{95}	34.0 Gy	25.7 Gy	22.7 Gy	0.003
V_{93}^{-2}	98.7%	98.7%	90.5%	< 0.001
Spinal cord		_		
D_{max}	25.7 Gy	19.3 Gy	23.3 Gy	0.003
$D_{1 cc}$	23.0 Gy	17.1 Gy	20.2 Gy	0.04
Brainstem				
D_{max}	28.2 Gy	22.3 Gy	24.9 Gy	0.007
$D_{1 cc}$	25.0 Gy	19.4 Gy	21.7 Gy	0.20
$\mathrm{D}_{1\%}$	26.1 Gy	20.2 Gy	22.9 Gy	0.12
Right parotid	(n = 12)			
D_{mean}	15.5 Gy	12.0 Gy	14.9 Gy	0.05
D_{50}	13.0 Gy	10.6 Gy	13.6 Gy	0.06
V ₂₆	44.6%	45.5%	55.5%	0.04
Left parotid				
D_{mean}	15.2 Gy	11.9 Gy	12.1 Gy	0.81
D_{50}	13.2 Gy	10.2 Gy	11.2 Gy	0.47
V ₂₆	45.2%	42.9%	42.2%	0.89
Mandible $(n = 9)$				
D_{max}	39.2 Gy	29.6 Gy	31.3 Gy	0.01
V ₆₀	11.0%	11.3%	18.2%	0.08
V ₇₀	0.04%	0.05%	4.5%	0.32

Abbreviations: PTV_{GTV} PTV_{CTV} = planning target volumes of gross tumor volume and clinical tumor volume, respectively; D_{max} = maximum dose; D_{99} = dose to 99% of the volume; D_{95} = dose to 95% of the volume; V_{93} = percent of volume receiving \geq 93% of the prescribed dose; $D_{1\ cc}$ = dose to 1 cc of the volume; $D_{1\%}$ = dose to 1% of the volume; D_{mean} = mean dose; D_{50} = dose to 50% of the volume; V_{26} , V_{60} , and V_{70} = percent of volume receiving \geq 26 Gy, \geq 60 Gy, and \geq 70 Gy, respectively.

Assessed impact on OAR doses not target dose

Contoured OARs on CBCTs and recalc'd with correction for HU differences

Clinical Investigation: Head-and-Neck Cancer

Monitoring Dosimetric Impact of Weight Loss ——With Kilovoltage (KV) Cone Beam CT (CBCT) During Parotid-Sparing IMRT and Concurrent Chemotherapy

Kean Fatt Ho, F.R.C.R.,* Tom Marchant, Ph.D.,† Chris Moore, Ph.D.,† Gareth Webster, Ph.D.,† Carl Rowbottom, Ph.D.,† Hazel Penington, B.Sc.,‡ Lip Lee, F.R.C.R.,§ Beng Yap, F.R.C.R.,§ Andrew Sykes, F.R.C.R.,§ and Nick Slevin, F.R.C.R.§

From *Academic Radiation Oncology, †North Western Medical Physics, ‡Wade Radiotherapy Research Centre, and §Department of Clinical Oncology, The Christie NHS Foundation Trust, Manchester, UK

Received Oct 10, 2010. Accepted for publication Jul 6, 2011

Weight loss and parotid shrinking did occur, but insignificant impact on OAR doses Results inconsistent with previous studies Impact of neoadjuvant therapy?

Demonstrates the benefit of 3D imaging
Discusses options of dose calculation from CBCT

Where did this weight loss occur?



Tumour Shrinkage and Weight Loss

- A lot of literature!!!
- Every patient is individual
 - > RTTs treat them and can see these subtle changes
- Dosimetric (and clinical) impact will depend on original DVH results
- Without 3D imaging, you cannot accurately visualise or account for this
- "The dosimetric impact of anatomic changes during radiotherapy was of lesser importance than the effects of IGRT repositioning" (Graff et al., 2012)

What Else? Variation in Shoulder Position

- The shoulders move independently from the isocentre
- This shoulder motion changes the path length of the beam
- Superior shoulder shift results in target coverage loss

Table 3 Target coverage in the C6-C7 region

	IMRT			VMAT		
	100%	98%	95%	100%	98%	95%
C6-C7						
No shift	97	98	100	94	97	99
5 mm superior	90	98	100	84	96	99
15 mm superior	23	53	94	16	35	72
C7-T2						
No shift	98	100	100	_	_	_
15 mm posterior	89	99	100	_	_	_

Percentage of the clinical target volume (CTV) in the C6-C7 region covered by the 100%, 98%, and 95% isodose lines with no shift and with superior shifts for IMRT and VMAT plans, as well as the percent coverage of the CTV in the C7-T2 region with no shift and a 15 mm posterior shift. All percentages were evaluated for Patient 1.



What Else? Variation in Shoulder Position

- This positional variation cannot be corrected with translational or rotational corrections
- This variation also caused an increase in OAR dose
 - ➤ Brachial Plexus increased by up to 7.2Gy

• In the absence of CBCT the angle of clavicle on AP EPI



Take Home Message

- "Complex and multifactorial dosimetric variations occur during head and neck IMRT." (Graff et al., 2012)
- Take caution due to tight margins, conformal techniques and proximity of radiosensitive structures
- Have an understanding of dosimetric impact of weight loss and shoulder motion
- Appropriate immobilisation is key. IGRT may help in assessment of this, but can not always correct for this.
- Recommend clear protocols to mandate imaging frequency and match structures

 ESTRO

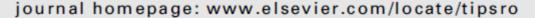
RTTs! If you are going to read one head and neck paper this year... Let it be this one!

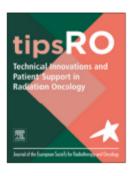
Technical Innovations & Patient Support in Radiation Oncology 1 (2017) 1-7



Contents lists available at ScienceDirect

Technical Innovations & Patient Support in Radiation Oncology





Practice guidelines

ESTRO ACROP guidelines for positioning, immobilisation and position verification of head and neck patients for radiation therapists



Michelle Leech ^{a,*}, Mary Coffey ^a, Mirjam Mast ^b, Filipe Moura ^c, Andreas Osztavics ^d, Danilo Pasini ^e, Aude Vaandering ^f

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c Hospital CUF Descobertas, Lisboa, Portugal

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e USCU Policlinico A. Gemelli, Rome, Italy

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Implementation of new protocols

Martijn Kamphuis MSc MBA Research Radiation Therapist IGRT

Department of Radiotherapy @AMC Amsterdam, the Netherlands



The Aim of the presentation

- Describing the process of implementing new technology and protocols
 - > Illustrated with different examples
- Sharing experience, tips and tricks



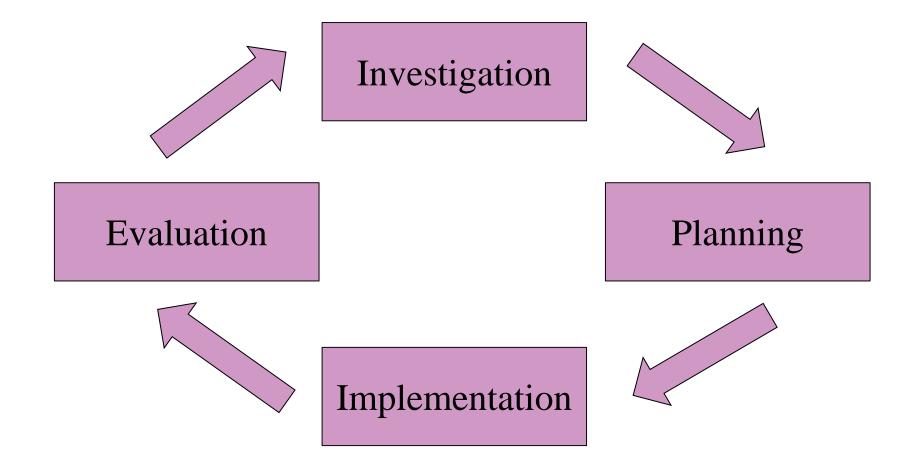
Have you ever been involved in clinical implementations of new protocols?

A. Yes

B. No



Protocol implementation





Preparation phase

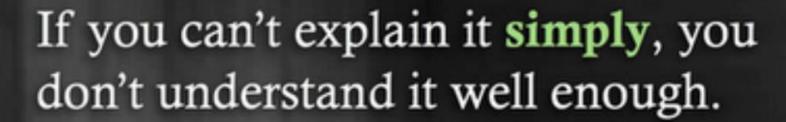
- Defining your goal, make sure it's clear
 - E.g. Implementing hypofractioned radiotherapy in your department for stage one and two lung cancer
 - Implementing adaptive radiotherapy for bladder cancer using a library of plans
- Creating a multidisciplinary team
 - Include all stakeholders that will get involved
 - Not only MD, physicist and RTT, but also manager, technicians
 - Define roles of individual team members
 - Who is doing what?

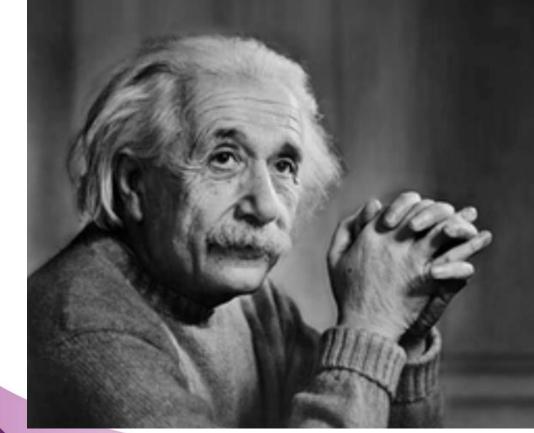


Preparation phase: Investigate!

- Literature reading
 - > Articles
 - Guidelines e.g. AAPM
- Join trails if possible
- Visit other institutes:
 - ➤ Learn from other ones' experience (and then do it better ⑤)
- Follow vendor trainings/courses/workshops
- ESTRO
 - ➤ (Live) Courses
 - Dove (www.estro.org)
 - Technology transfer grant







Albert Einstein

Planning/protocol writing phase

- Organize multiple meetings to discus the protocol
 - Come to a shared vision
- Write a project plan/protocol
 - Define all tasks and responsibilities
 - Check weather task can be performed parallel
 - Create a timeline
 - Start from the deadline ©
 - Decisions have to been made, sooner or later
- Prospective Risk Analyses
 - Helps to think ahead



Implementation Phase

Critical conditions for proper implementation

- Treatment protocol should be:
 - > Well described and well defined task
 - Approved by staff
 - Available for everyone



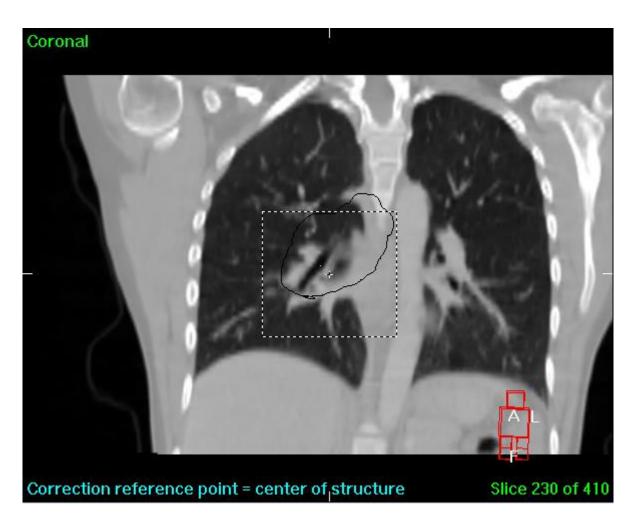
Well described and well defined tasks

Example: protocol for dealing with anatomical changes

- In room imaging started off as a single check between CT and Linac
- Nowadays we are more aware of anatomical changes



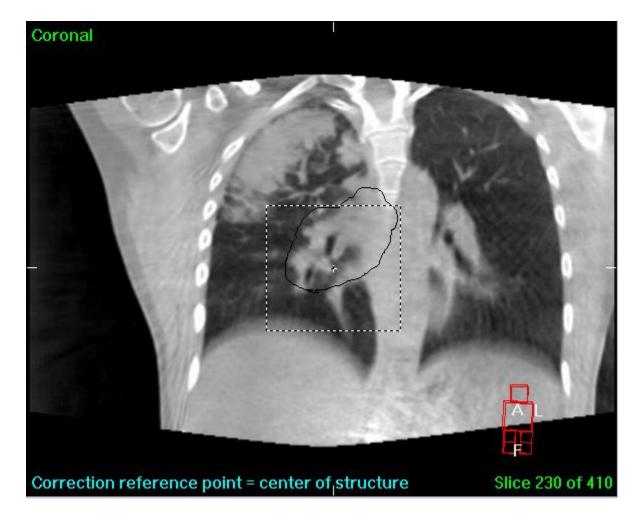
Anatomical changes



Ref-CT



CBCT







CBCT
Ref-CT



Slice 138 of 264 Transverse Scan Time: 19.08.2009 09:19:23.000 09.2009 15:04:20.671

CBCT

Ref-CT



Atelectases

- Dosimetry changes due to atelectases
- Tumor position changed
- Rescanning/replanning necessary

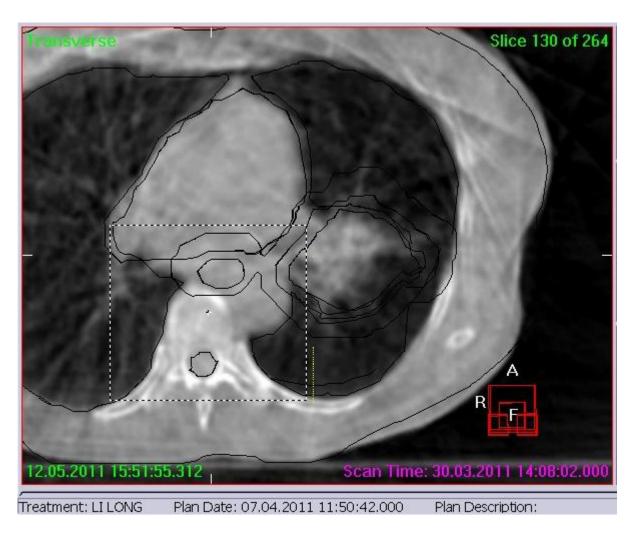




CBCT
Ref-CT



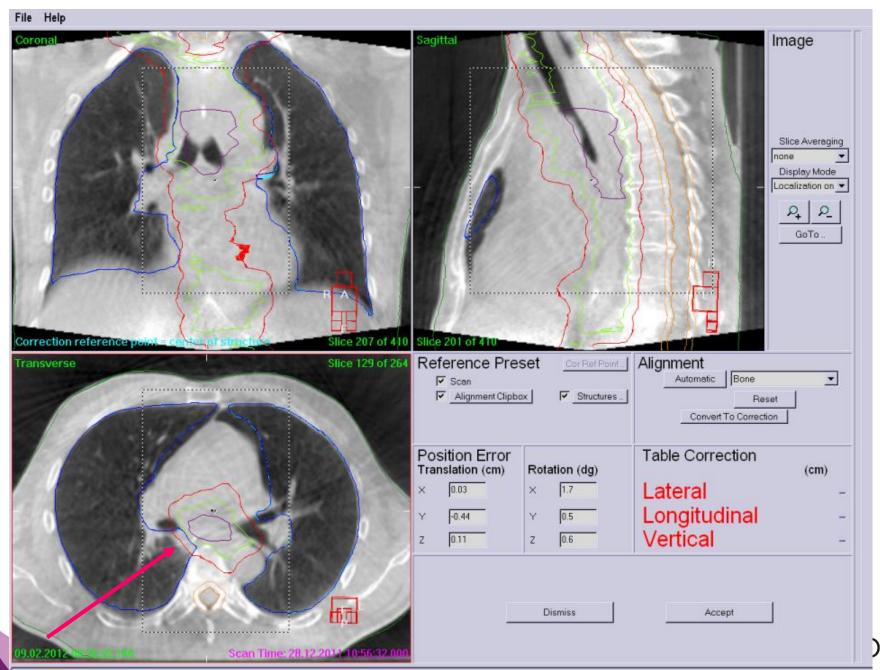
Tumor regression





Tumor shift

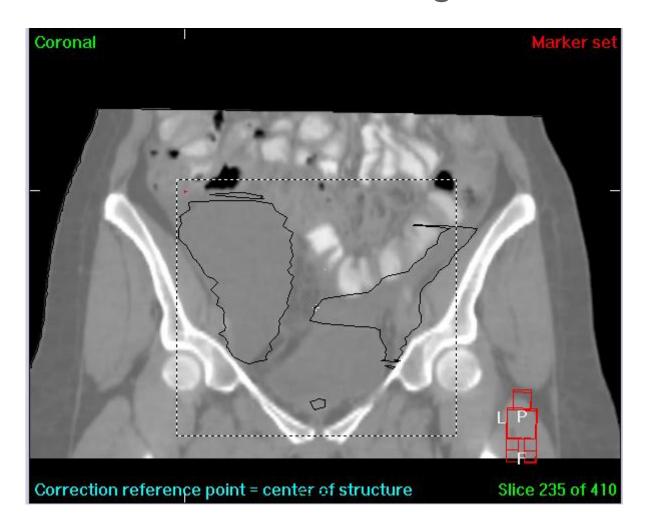




Rare changes

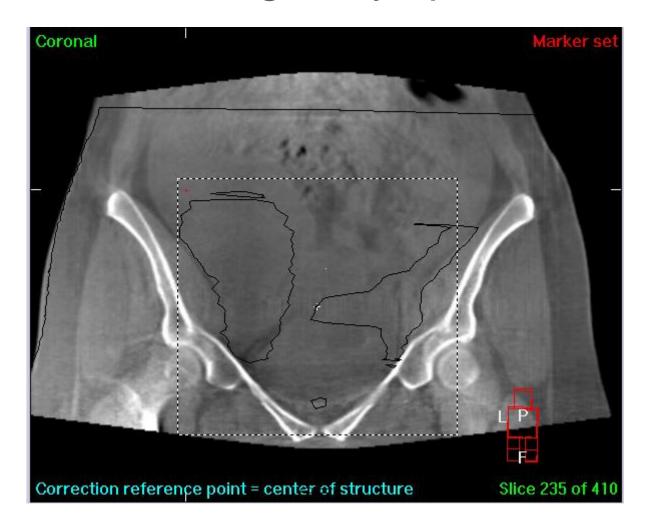


Rare changes





Change in lymphocele





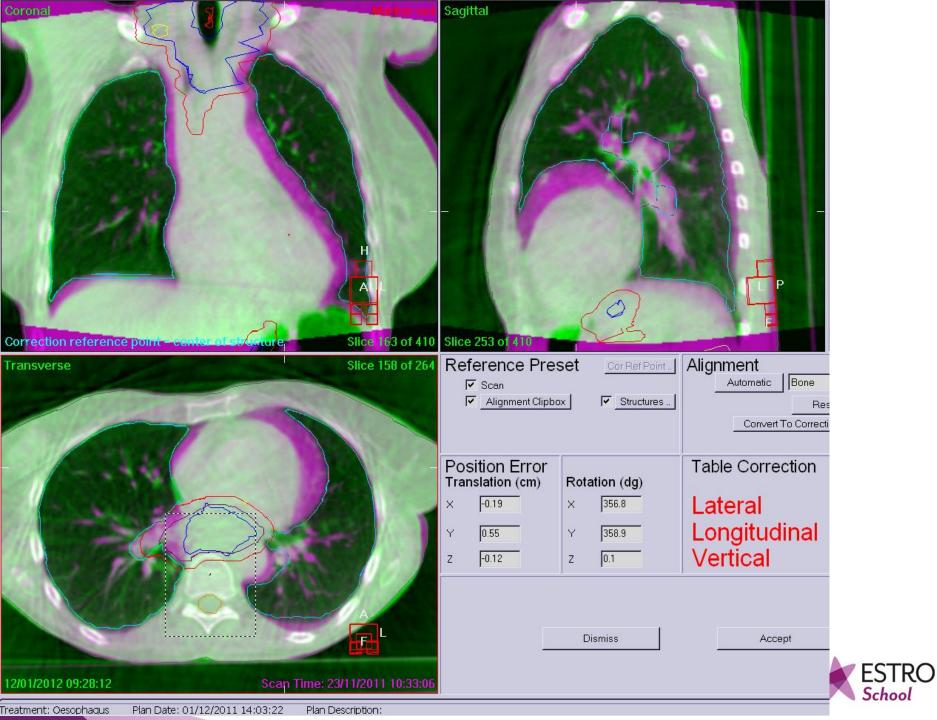
Weight loss



File Help Coronal Sagittal **Image** Slice Averaging none Display Mode Green-purple 🔻 GoTo. Correction reference point = center of structure Slice 273 of 410 Slice 197 of 410 Reference Preset Alignment Slice 177 of 264 Transverse Cor Ref Point Bone Automatic ▼ Scan ✓ Alignment Clipbox Structures. Choose method of alignm Convert To Correction Position Error **Table Correction** Translation (cm) Rotation (dg) (cm) -0.05 2.1 Lateral Longitudinal -0.43 0.0 Y Vertical -0.25 1.9 Z Dismiss Accept 01.12.2011 14:20:11.671 Scan Time: 02.11.2011 10:22:24.000

Cardiac changes





Summary

- Anatomy is changing during treatment
- RTT is the person most likely to detect
 - > Should be her/his responsibility
- You can't bother the doctor or physicist with everything...



How to deal with changing anatomy?*

- Call doctor before treatment
 - > Change in atelectases
 - ➤ GTV and/or CTV outside PTV
- Contact doctor that day or the day after
 - Mild tumor progression
 - Tumor regression
- Contour changes (physicist)
 - >2 cm
 - > > 1cm H&N and extremities

*Inspired by:

INTRA THORACIC ANATOMICAL CHANGES FOR LUNG CANCER PATIENTS DURING THE COURSE OF IRRADIATION: HOW TO RESPOND?

S. Conijn¹, J. Belderbos¹, J. Knegjens¹, M. Rossi¹, J. J. Sonke¹, P. Remeijer¹

The protocol...

- Describes were to look at
- Describes what do
- Describes who to contact
- Describes at what speed actions have to take place



Implementation Phase

Critical conditions for proper implementation

- Treatment protocol should be:
 - Well described and tasks well defined
 - Approved by staff
 - Available for everyone
- Education and training of professionals:
 - Really depends on subject
 - Preferable as practical as possible
- Example: *Bladder ART*



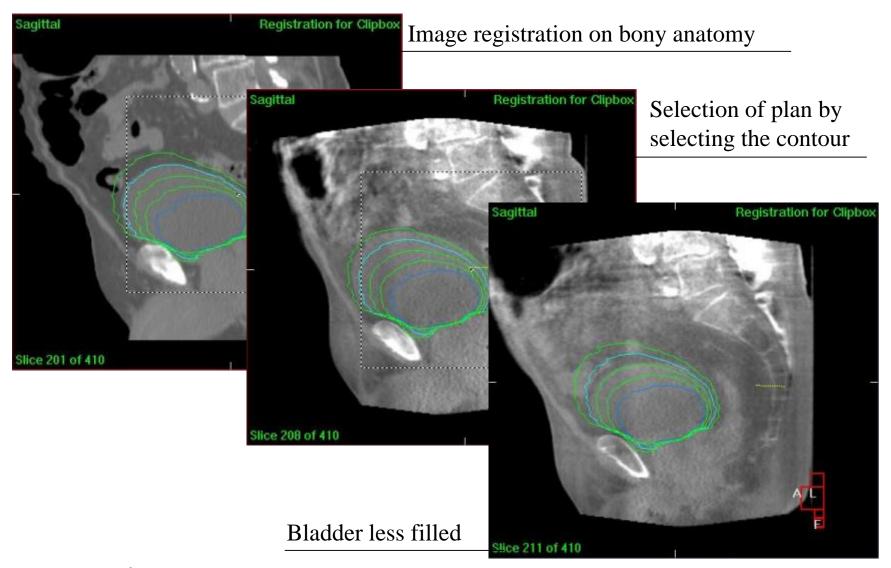
Plan of the day

Inter- and extrapolation of bladder contours

5 plans are generated on the TPS (Oncentra, Elekta)

Images: Jorrit Visser





Images: Rianne de Jong



Demo database ART blaas

4 patients with two reference CT

82 Conebeam CT-scans

5 structures per patient/scan:

$$0 - 33 - 67 - 100 - 133\%$$

12 observers

Interobserver study:

1^e measurement workshop 2^e measurement



Implementation Phase

Critical conditions for proper implementation

- Treatment protocol should be:
 - > Well described and well defined task
 - Approved by staff
 - Available for everyone
- Education and training of professionals:
 - Really depends on subject
- Implementation date
 - Properly communicated
 - Repeat communication just before start
- Use a predefined checklist



Evaluation phase

- Phase that ignored often
 - The work just started....
- Space to correct for mistakes
- Evaluate
 - Ask for feedback from your colleagues:
 - Pro active: create a feedback session
 - Data to validate new procedure
 - publish
 - Monitor your processes



Bladder ART: Safety-net plan selection

1^e week

- Doctor, physicist and IGART RTT on the linac
- Fixed moment

Starting of the 2^e week

- ➢ IGART RTT on the linac
- Fixed moment

After 10 patients

evaluation and feedback - database oefenpatiënten? -

Once weekly one dedicated IGART RTT check all decisions:

- Was the "right" selected?
- o Is the used model still valid?
- o Was the used treatment plan in the R&V system selected?



Conclusions

Implementing protocols:

- Investigate
- Plan
- Implement
- Evaluate

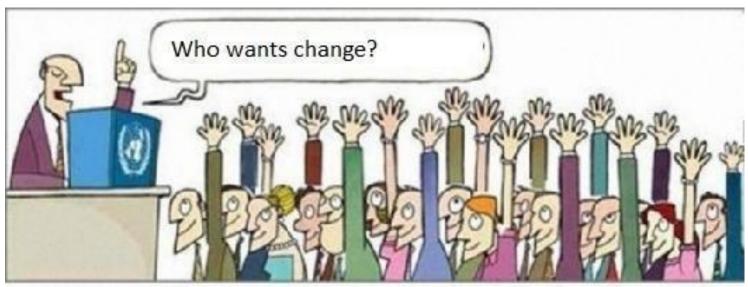
Implementing protocols is a change process:



John Kotter's 8 steps of change









ESTRO School

Thank you for your attention!



Who is doing what in Radiation Therapy

Rianne de Jong *RTT*, Amsterdam Medical Centre



m.a.j.dejong@amc.uva.nl



Survey

Questionnaires to participants of ESTRO course on "IGRT in clinical practice" in 2006-2010:

48 hospitals

19 countries

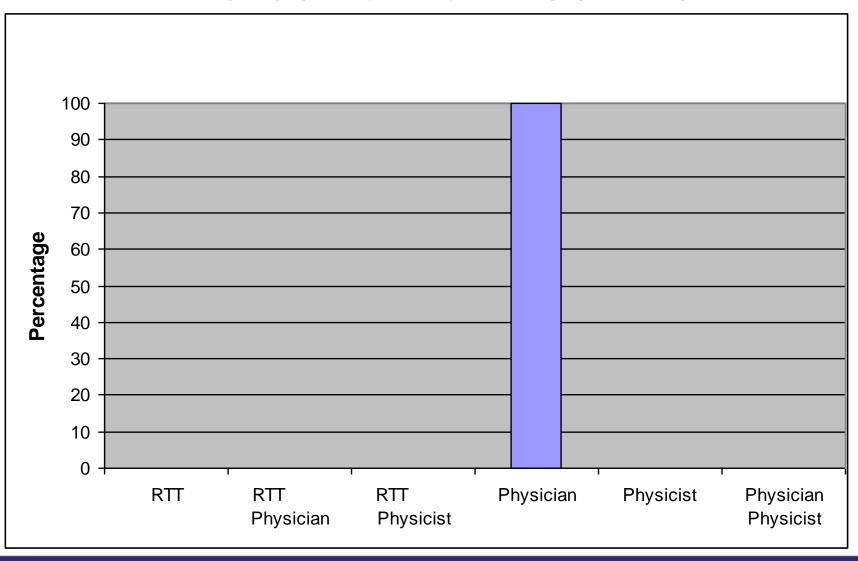
Survey

- 1. Indication/Design of Radiation Treatment
- 2. Pre treatment imaging: CT/simulation
- 3. Delineation
- 4. Treatment Planning
- 5. Treatment
- 6. Image Guidance/Adaptation treatment



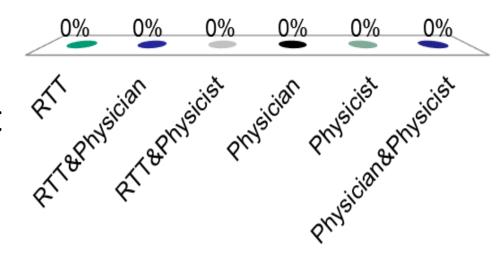
- Radiation Therapy Technicians (RTT)
- Physicians
- Physicists

1. Indication of treatment

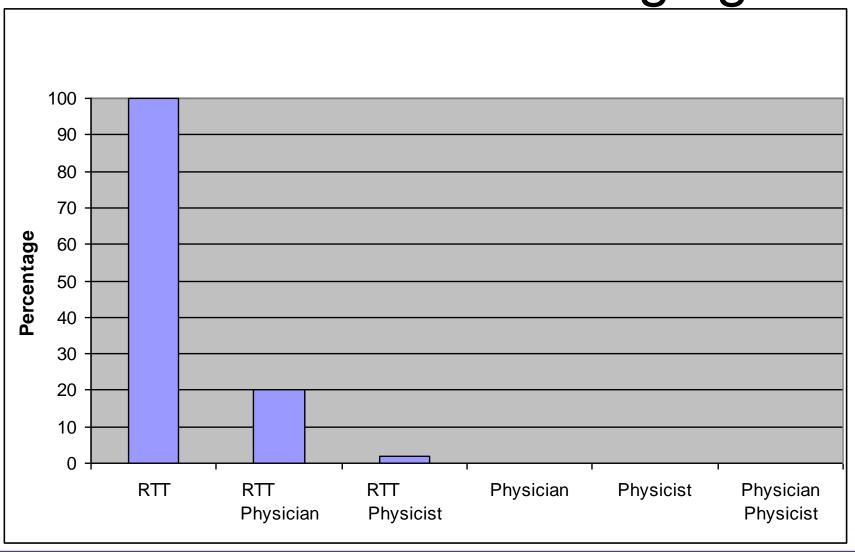


1. Indication of treatment

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist

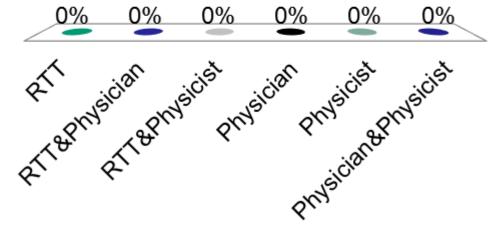


2. Pre-treatment Imaging

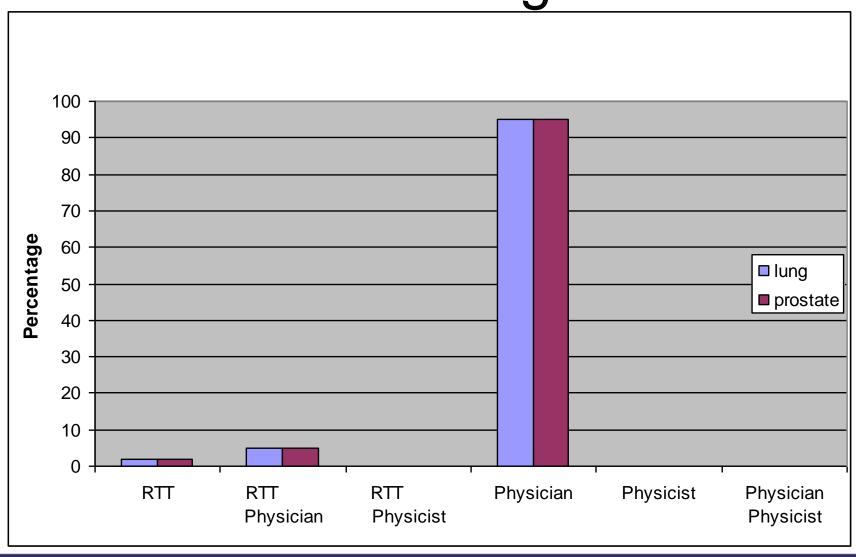


2. Pre treatment Imaging

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist

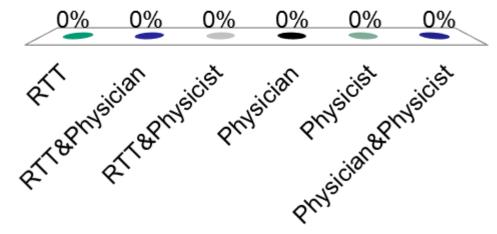


3. Delineation: Target Volume

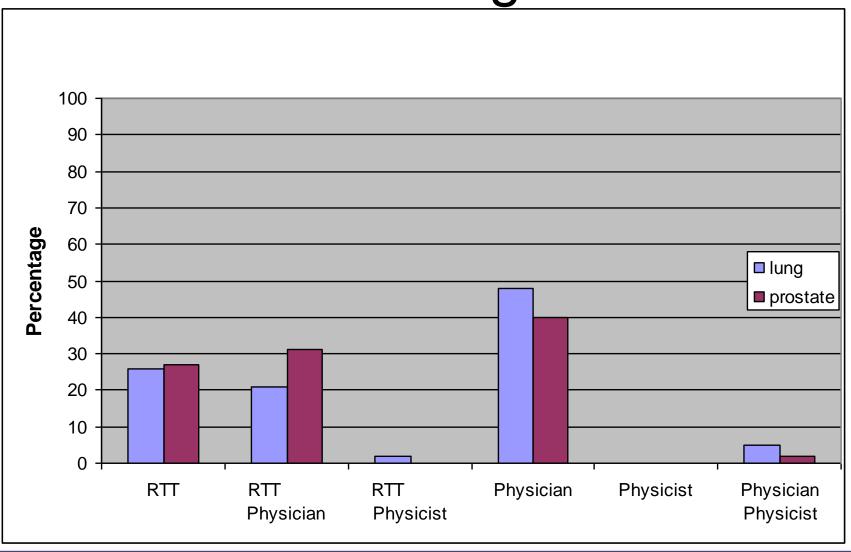


3. Delineation Target Volume

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist

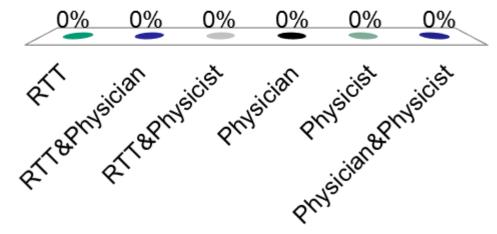


3. Delineation: Organs at Risk

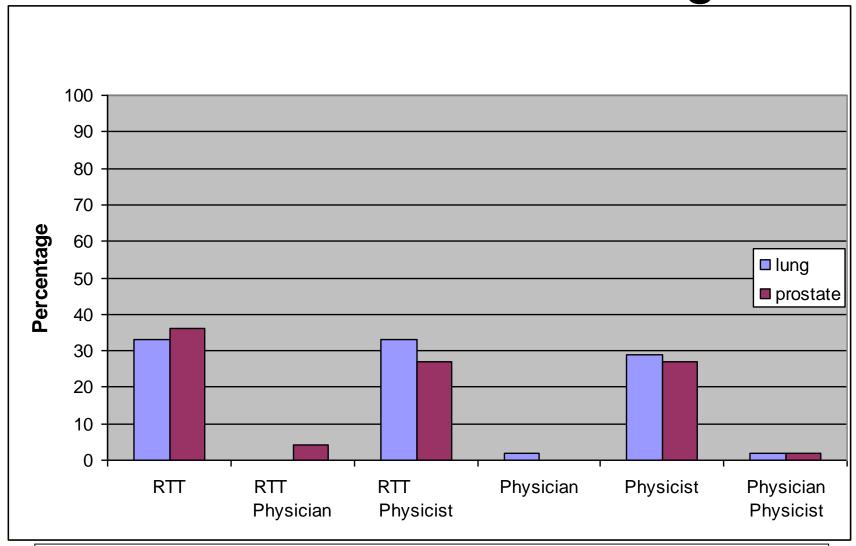


3. Delineation Organs at Risk

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist



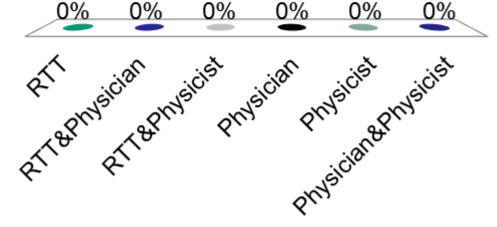
4. Treatment Planning



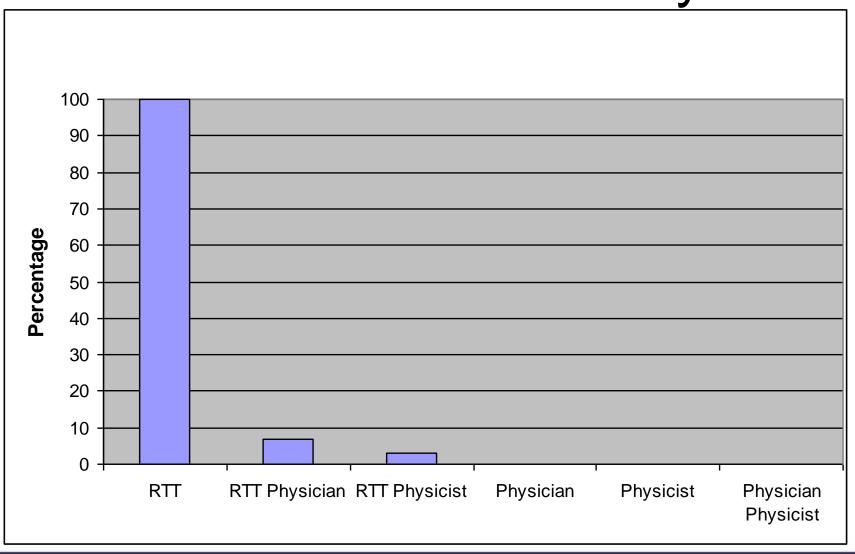
RTT: supervised and/or accepted by physician or physicist

4. Treatment Planning

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist

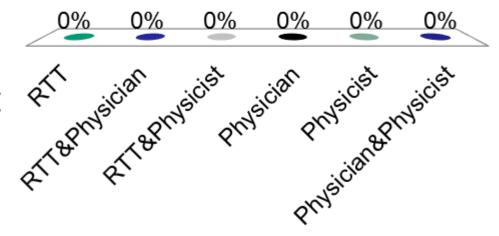


5. Treatment Delivery

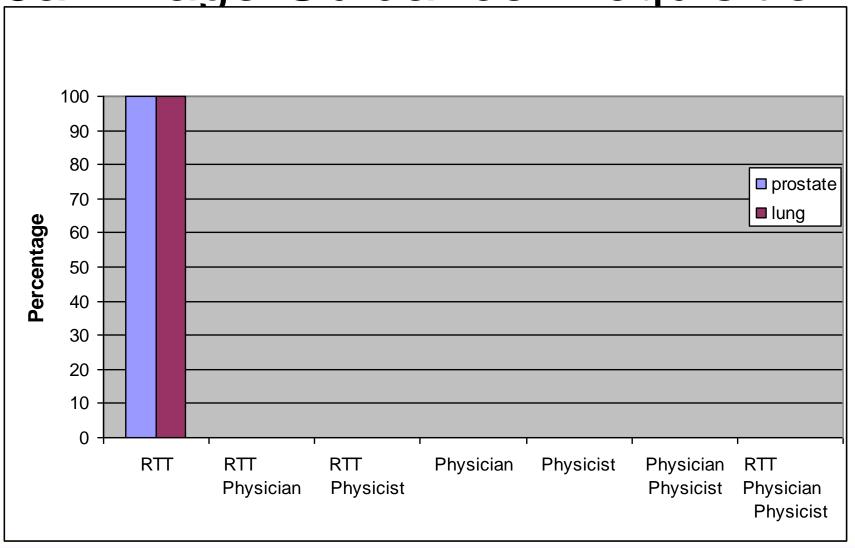


5. Treatment Delivery

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist

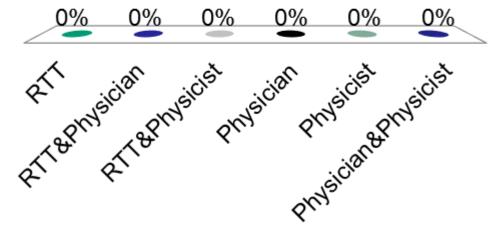


6a. Image Guidance: Acquisition

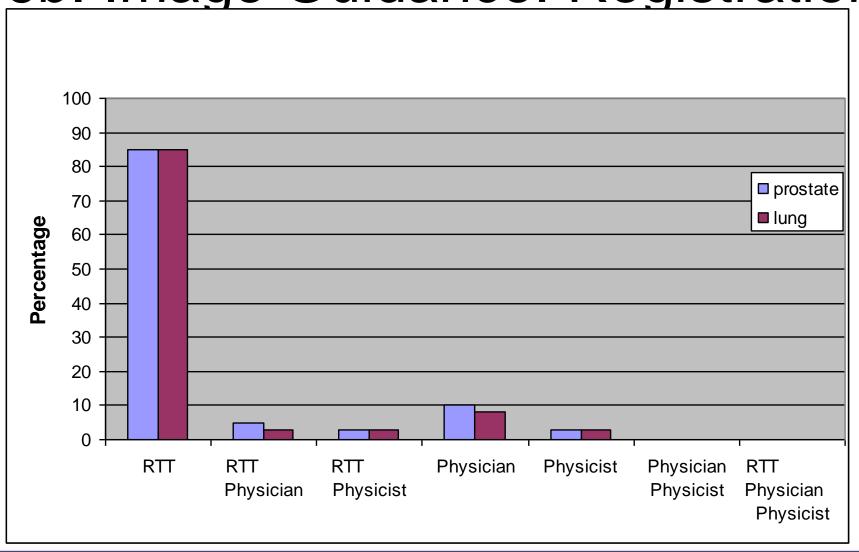


6a. Image guidance: Acquisition

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist

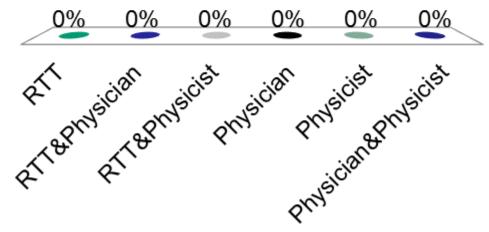


6b. Image Guidance: Registration

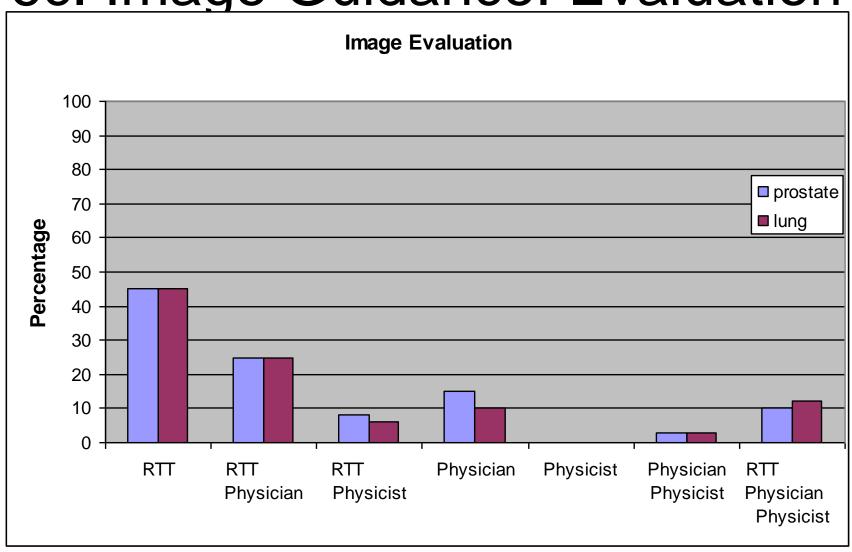


6b. Image Guidance: Registration

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist

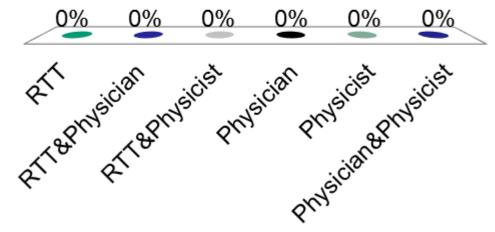


6c. Image Guidance: Evaluation



6c. Image Guidance: Evaluation

- A. RTT
- B. RTT&Physician
- C. RTT&Physicist
- D. Physician
- E. Physicist
- F. Physician&Physicist



Who is doing what?

Conclusion: Largest differences in *Treatment Planning* and *Image Guidance*.

Why? What are the variables in the different departments that could have an influence on these differences?

- RTT education / training
- Department size
- Resources per treatment machine
- IGRT modalities
- Culture / History
- Money

RTT training / Education

Majority:

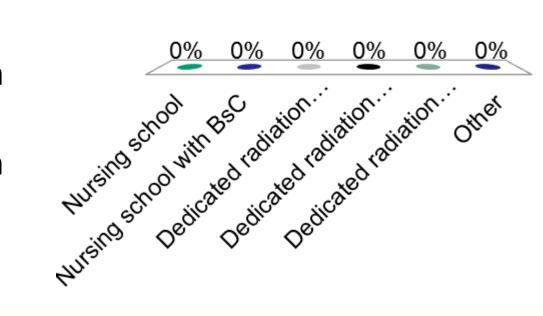
- 3 years of classroom combined with clinical intern hours
 - → bachelor degree

Also:

- 3 years of nursing school with bachelor degree with additional theoretical or clinical RTT training ~1 year.

Training & Education

- A. Nursing school
- B. Nursing school with BsC
- C. Dedicated radiation therapy
- D. Dedicated radiation therapy with Bsc
- E. Dedicated radiation therapy with MsC
- F. Other



RTT training / Education

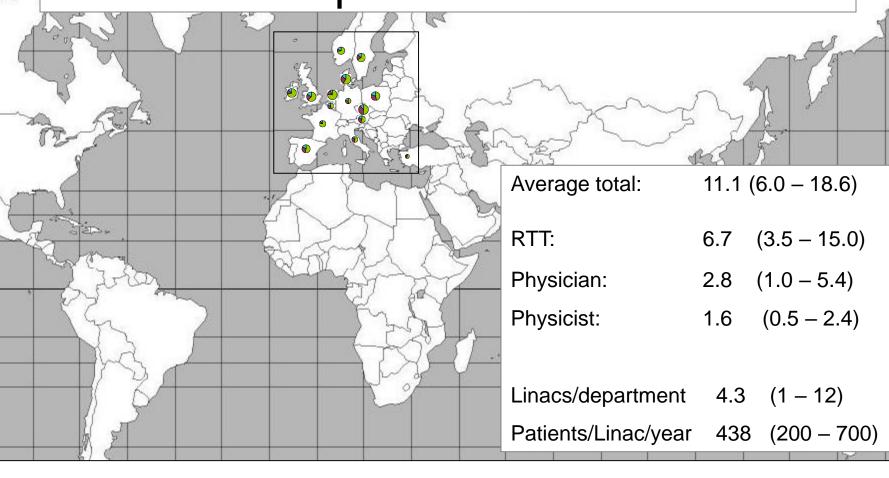
Majority:

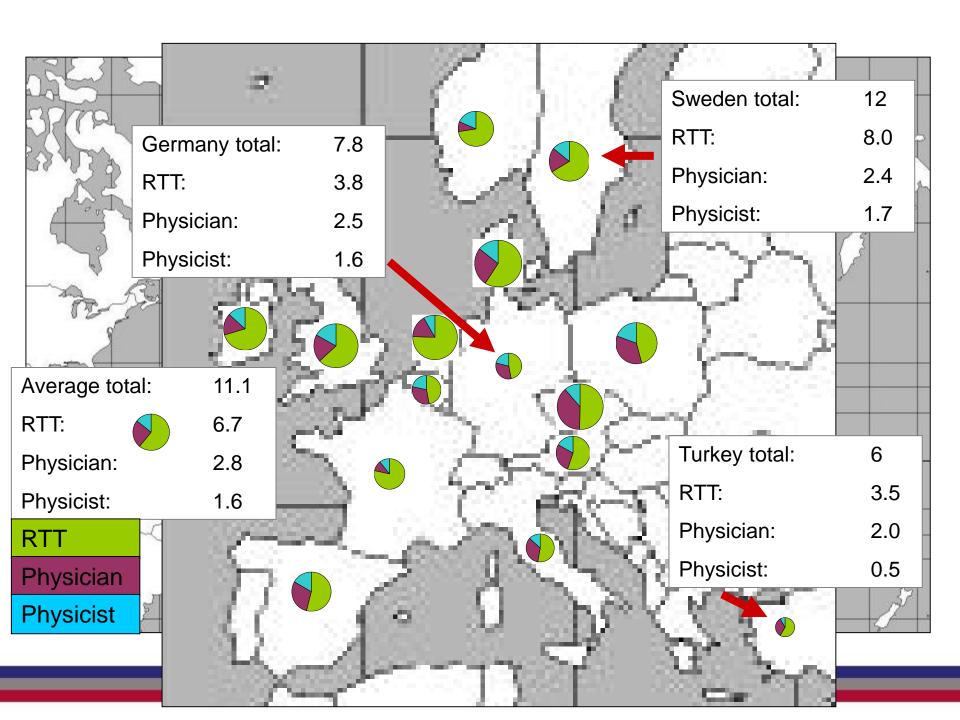
- 3 years of classroom combined with clinical intern hours
 - bachelor degree

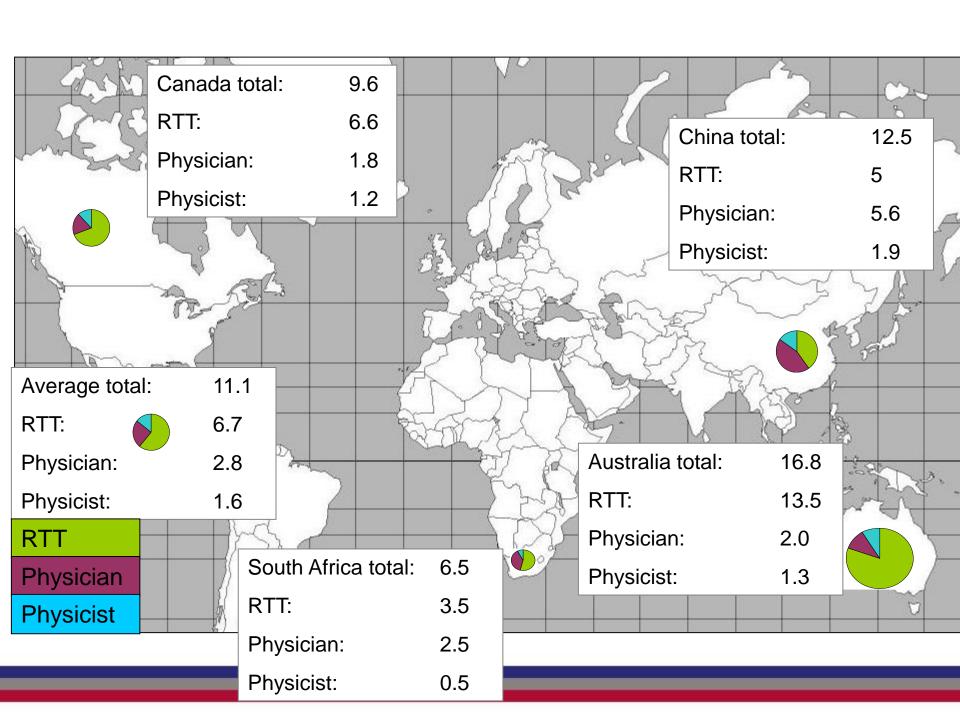
Also: Does not correlate • 2 or 4 years or classroom combined with climical in

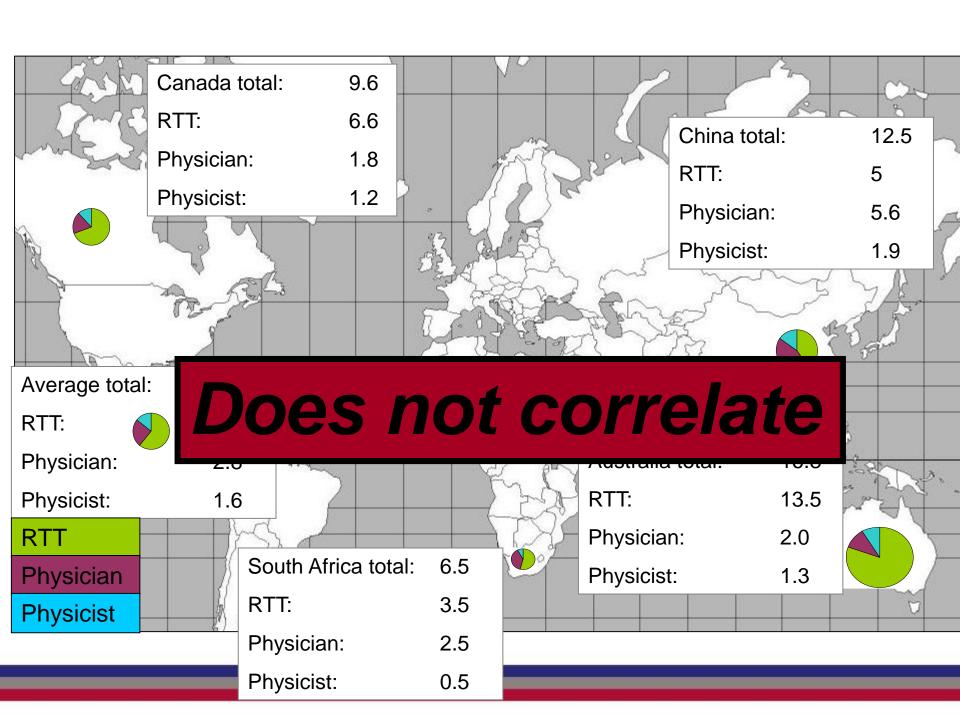
- hours bachelor degree
- 3 years of nursing school with bachelor degree with additional theoretical or clinical RTT training ~1 year.











IGRT Modalities:

2D Portal Images	79%
------------------	-----

2D kV Images 6%

kV Conebeam CT 66%

MV Conebeam CT 17%

IGRT protocols are:

– 1	Tumor	site	specific	100%
------------	-------	------	----------	------

Patient specific 18%

– Physician specific2%

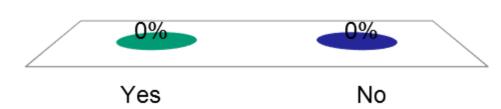
IGRT modalities: 2D MV

A. Yes



IGRT modalities: 2D kV

A. Yes



IGRT modalities: 3D kV

A. Yes



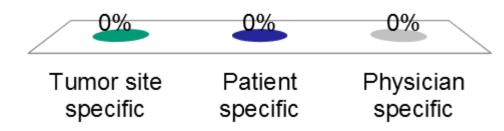
IGRT modalities: 3D MV

A. Yes

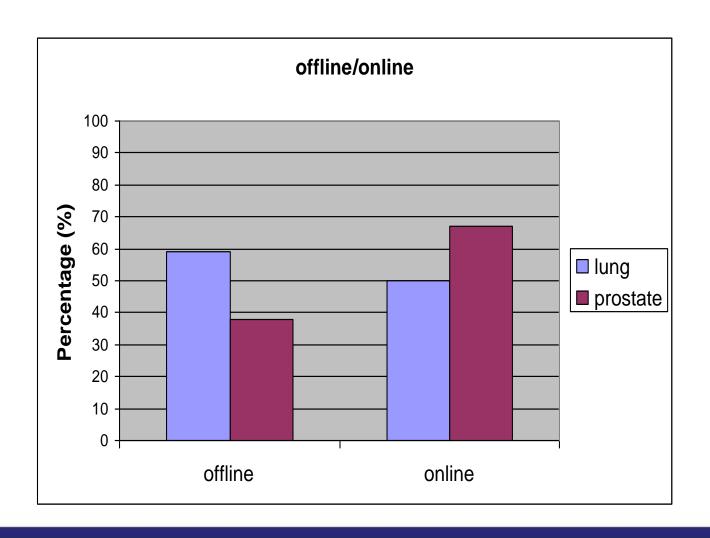


IGRT protocols are

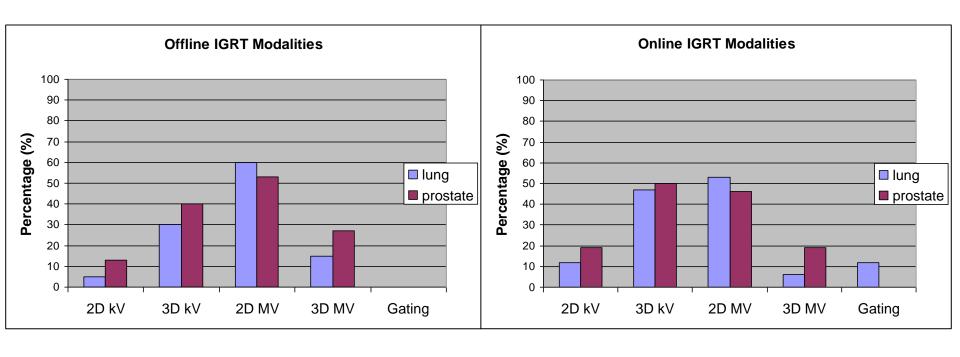
- A. Tumor site specific
- B. Patient specific
- C. Physician specific



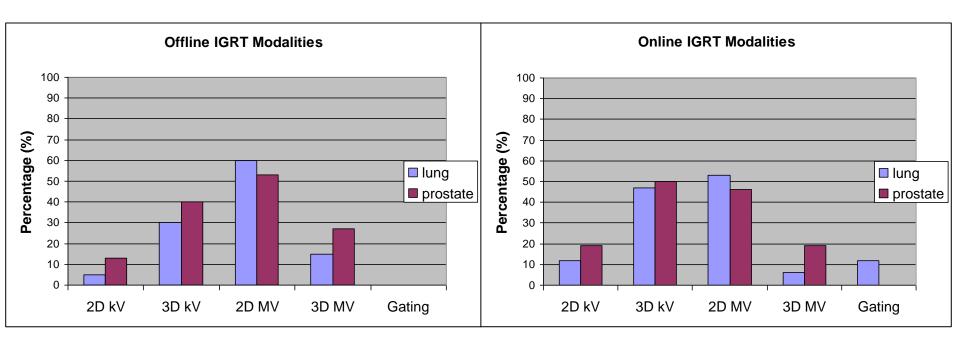
2D Portal Images 69% kV Conebeam CT 67% MV Conebeam CT 18%



2D Portal Images 69% kV Conebeam CT 67% MV Conebeam CT 18%



2D Portal Images 69% kV Conebeam CT 67% MV Conebeam CT 18%



→ Adaptive Radiation Therapy... 0%

2D Portal Images 69% kV Conebeam CT 67% MV Conebeam CT 18%



→ Adaptive Radiation Therapy... 0%

Who is doing ART?

A. Yes



Summary

Large variation between departments in:

- Amount of resources per linac
- Their distribution in different disciplines:
 - Treatment planning
 - IGRT evaluation

Some Variables

- RTT training and education
- Department size
- Resources per treatment machine
- IGRT Modalities

» Culture - History

» Money

Not decisive

Might consider different solutions?

Questions & Discussion



m.a.j.dejong@amc.uva.nl



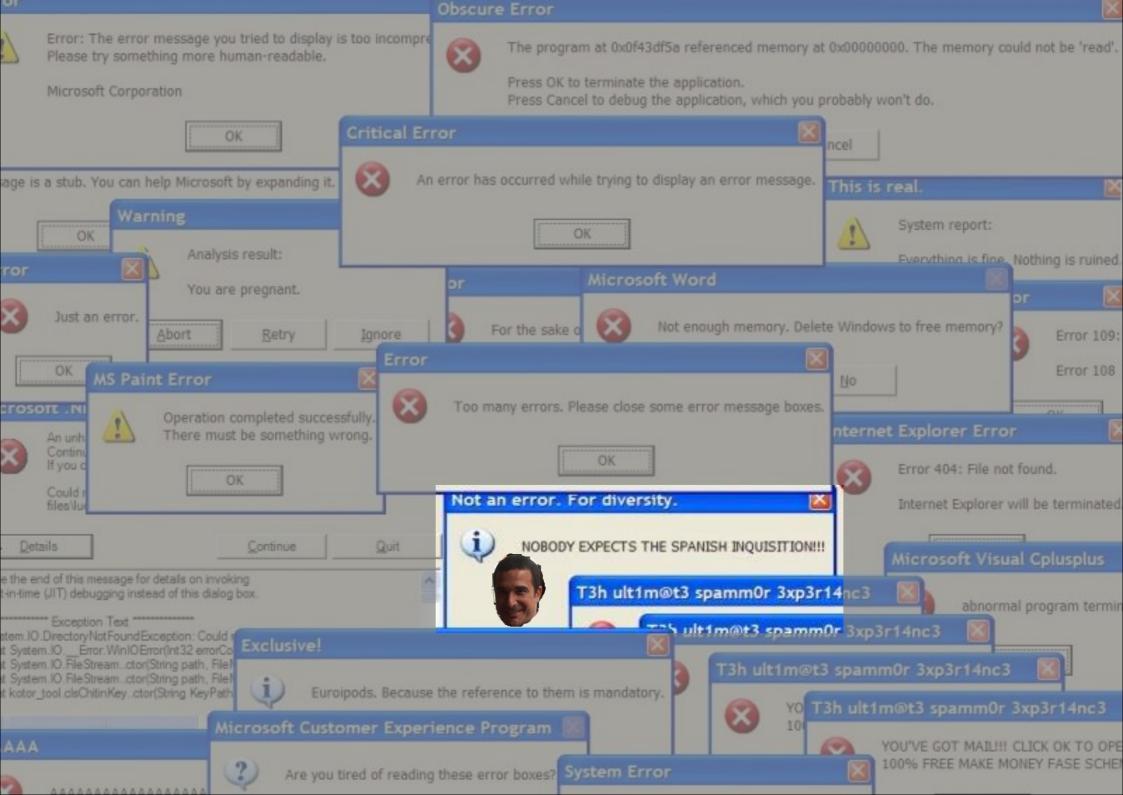


Error management

Peter Remeijer
Department of Radiation Oncology
The Netherlands Cancer Institute







More errors?

- Transfer errors (planning → linac)
- Linac errors (both dosimetric and geometric)
- Dosimetric errors in plan
- Input errors
- Patient setup (e.g. CT reference to isoc shifts)
- Select the right patient / treatment in all systems

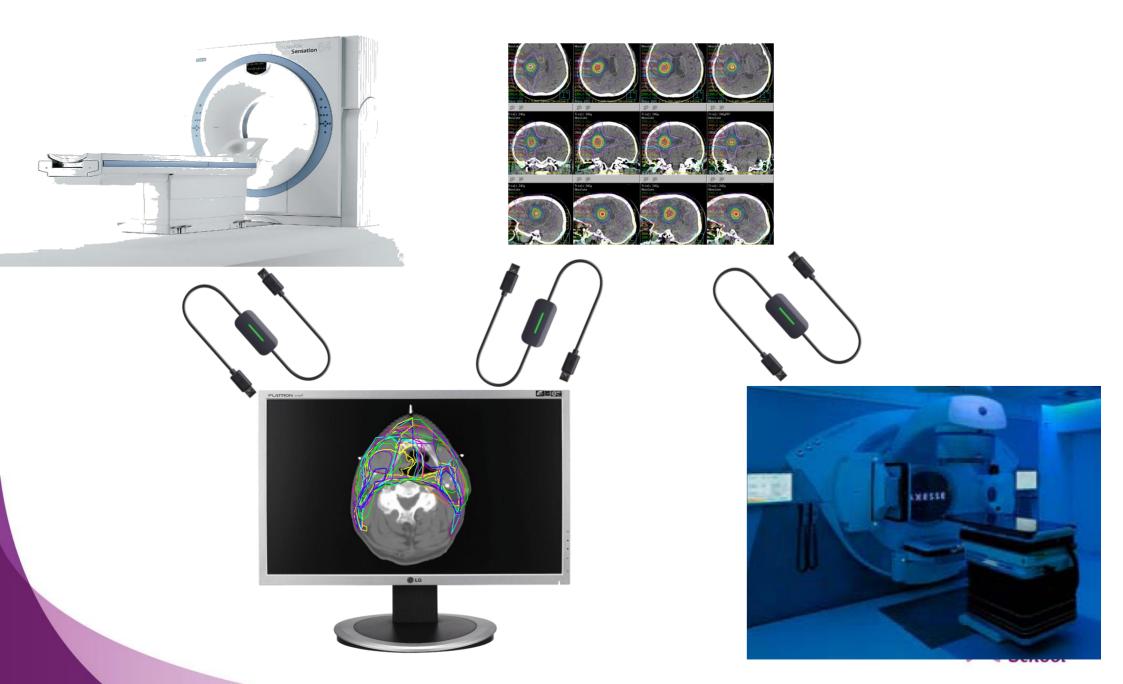


More errors?

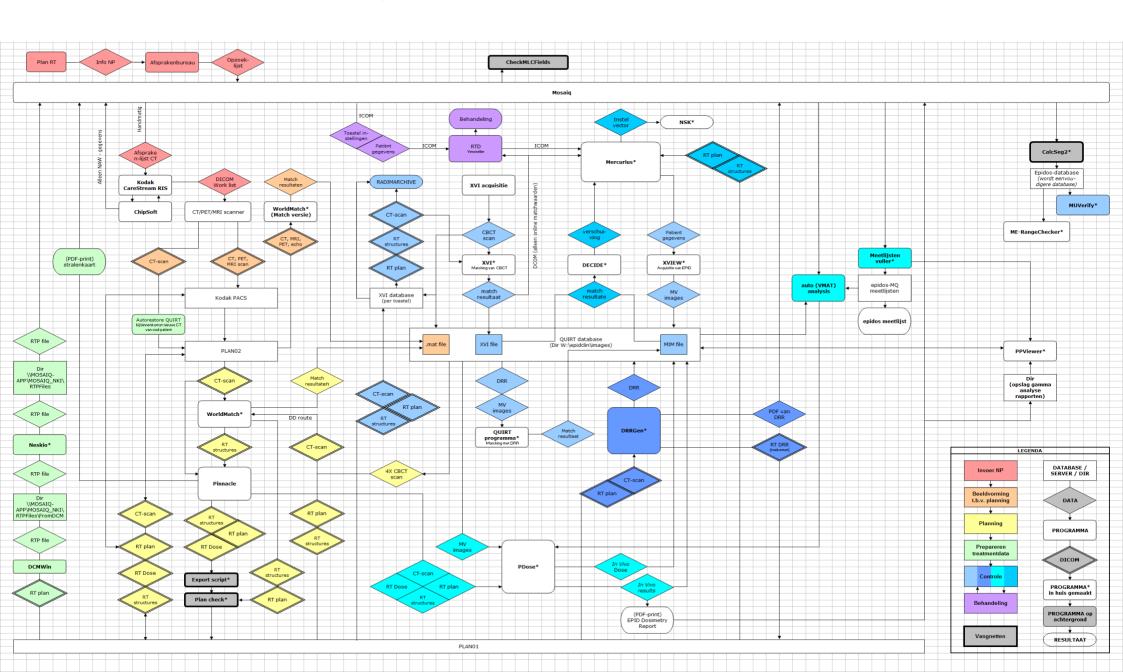
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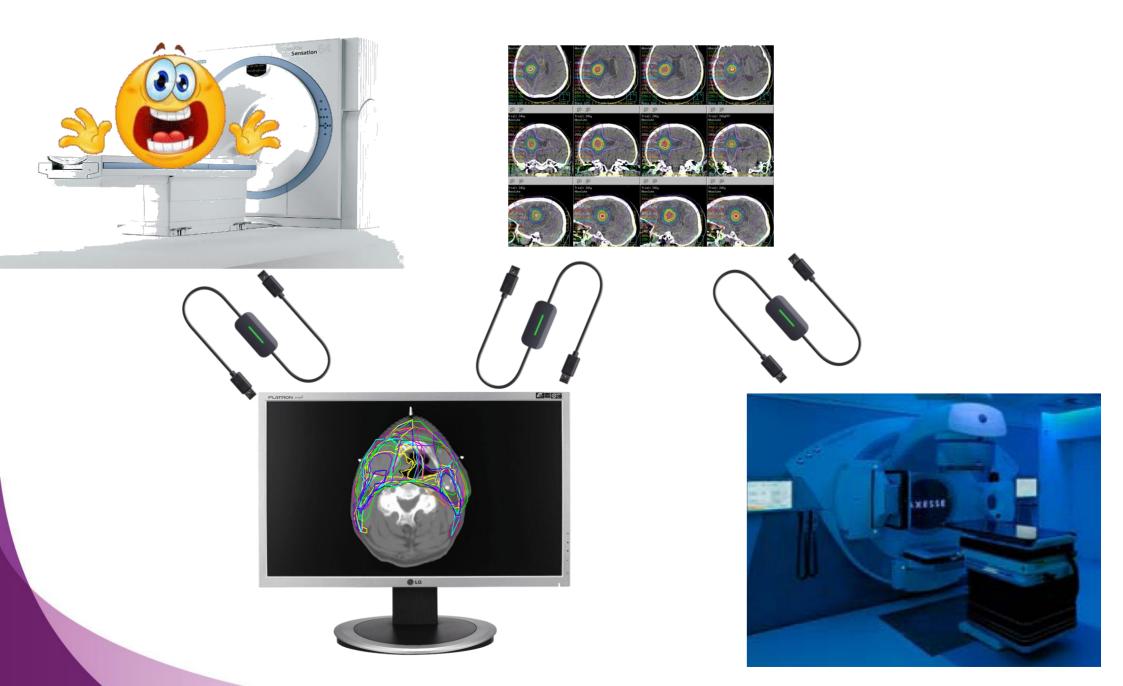
Errors and the radiotherapy "chain"



The radiotherapy "chain"



"Chain test" a.k.a regression test with phantom



Regression testing

- Run a phantom through the whole treatment chain and check for problems / errors
 - May be necessary to do this for different situations, i.e. HFS, HFP, etc
 - New methods, e.g. ART, library of plans, new planning techniques (VMAT)
- This will check
 - Connectivity
 - Systematic equipment and software errors
 - Overall dosimetry
 - Overall geometry



More errors?

- Transfer errors (planning → linac)
- Linac errors (both dosimetric and geometric)
- Dosimetric errors in plan
- Patient setup (e.g. CT reference to isoc shifts)
- Input errors
- Select the right patient / treatment in all systems



Independent MU checks

- Recalculates the dose, based on the plan parameters from the planning system (or v.v.)
- This will check (in theory)
 - Amount of monitor units
 - Problems with plan normalization
 - Computation errors of planning system
- Third party software
 - Lots of software around (small companies)
 - Check what it really checks
 - Test with intentional errors



MU range checking

- In house NKI development, but easy to build
- Plans following a certain protocol, e.g. prostate
 - Amount of MU for a VMAT plan will be similar for each patient
 - Depends a little on patient size, etc
 - → MU range check
 - If patient does not fall within the range, something may have gone wrong
 - Check by physics
 - About 5-10%
 - Usually anatomical reasons
 - Some errors found (wrong dose specification point, #fractions)

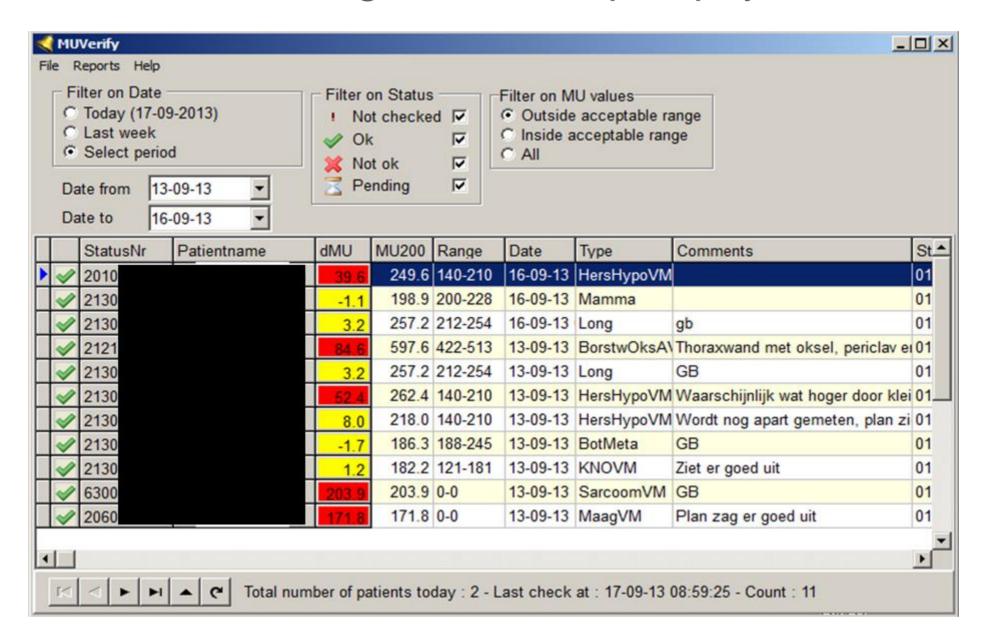


MU range checking

- Plan type depends on
 - Careplan name (brain, breast, prostate, etc)
 - RX-site name (plan name), e.g. Sacrum <231290>
 - Number of beams
 - Number of segments
 - Energy
 - Fraction dose
- Range for each type

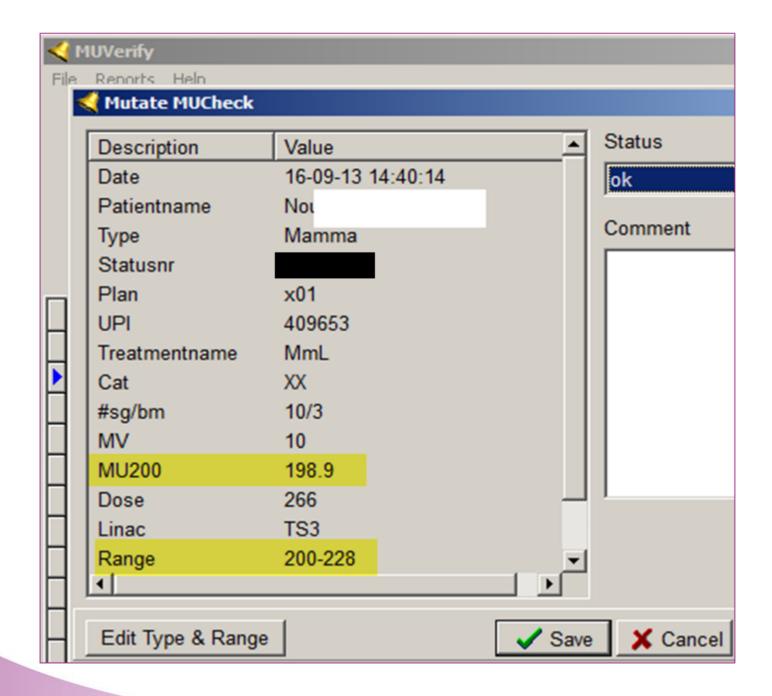
```
CP
      |Nbeam| Nsegm | Energy | Fr.Dosis
                                                      | Min
                                                              Max
                                              Type
                   70 | 6 | 10 |
                                 180 | 300
               2 |
                                                        188
                                                              261
Anus
                                             Anus
                  180 | 6 | 10 | 180
Blaas
        11 21
              70|
                                     | 400
                                             BlaasVM |
                                                        158
                                                              218
                   60 | 6 |
                            10 |
                                 180 | 800
Cervix| 2|10| 2|
                                              Gyn
                                                        221
                                                              284
```

Automated message on desktop of physicist





Automated message on desktop of physicist



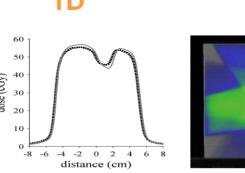


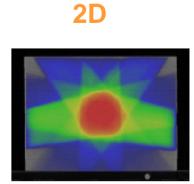
In-vivo portal dosimetry

in most centres today:

not 3D

(hg) sop 20 10





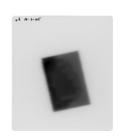
not in vivo





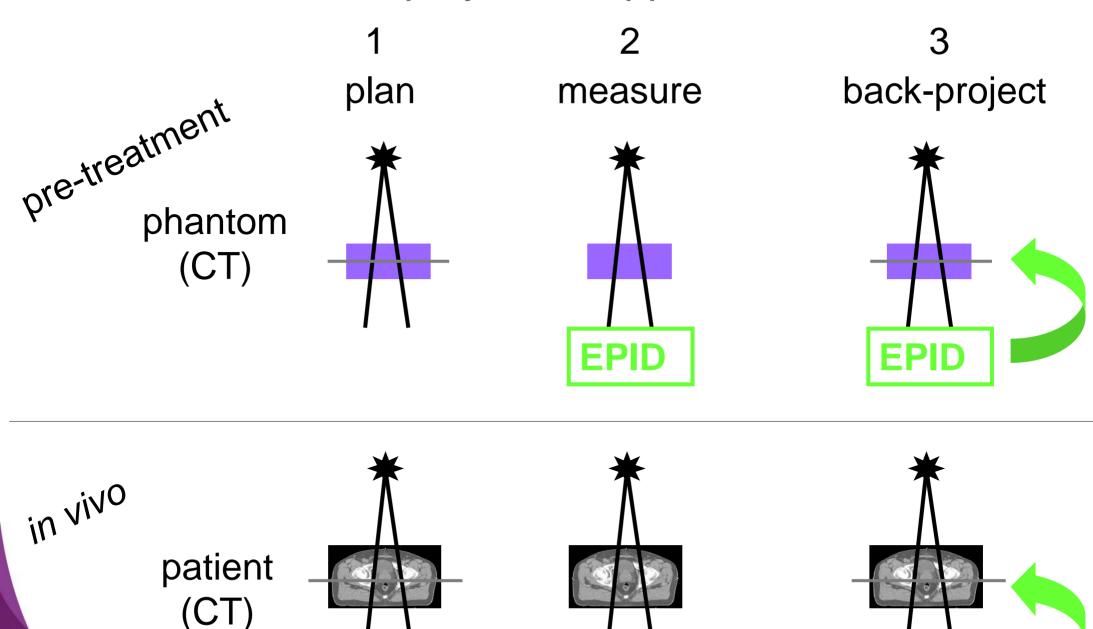
not with an EPID





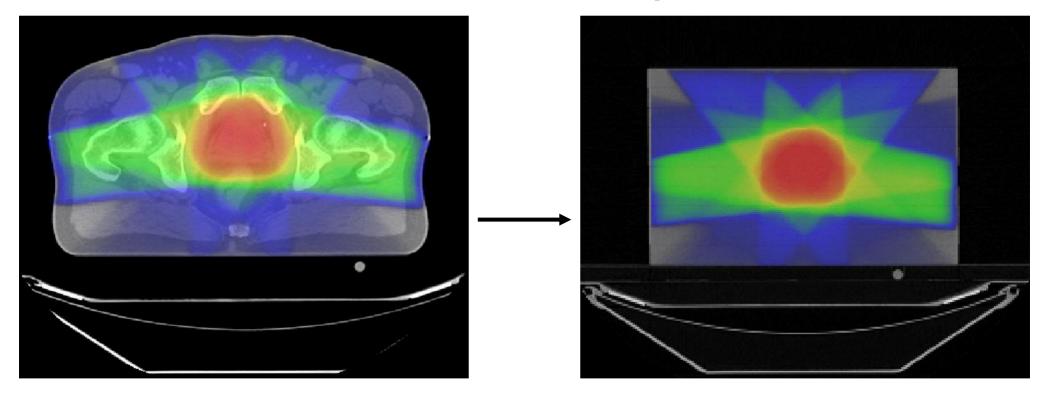


The NKI back-projection approach



EPID

Pre-treatment: in a phantom



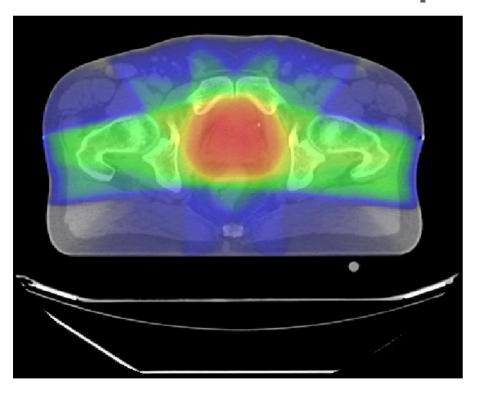
checks: plan deliverability

dose calculation

extra time: about 1 hour



In vivo: in the patient





checks: plan deliverability

dose calculation

anatomy changes

random delivery errors

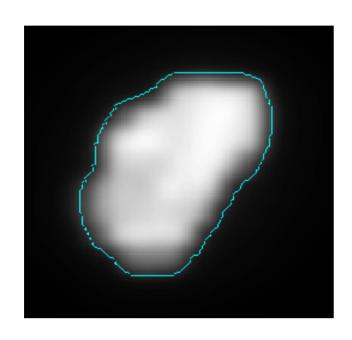
extra time: ~ 25 min in case of an error

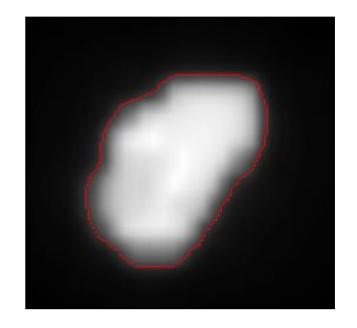
+ 30s/day



Field-by field reference vs calculated or measured dose

how do we compare them in 2D?



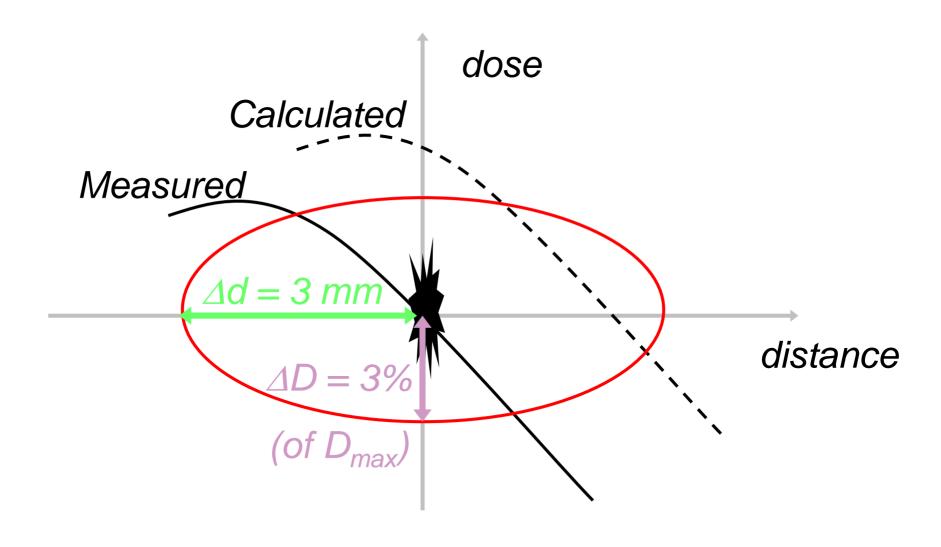


PLAN

EPID

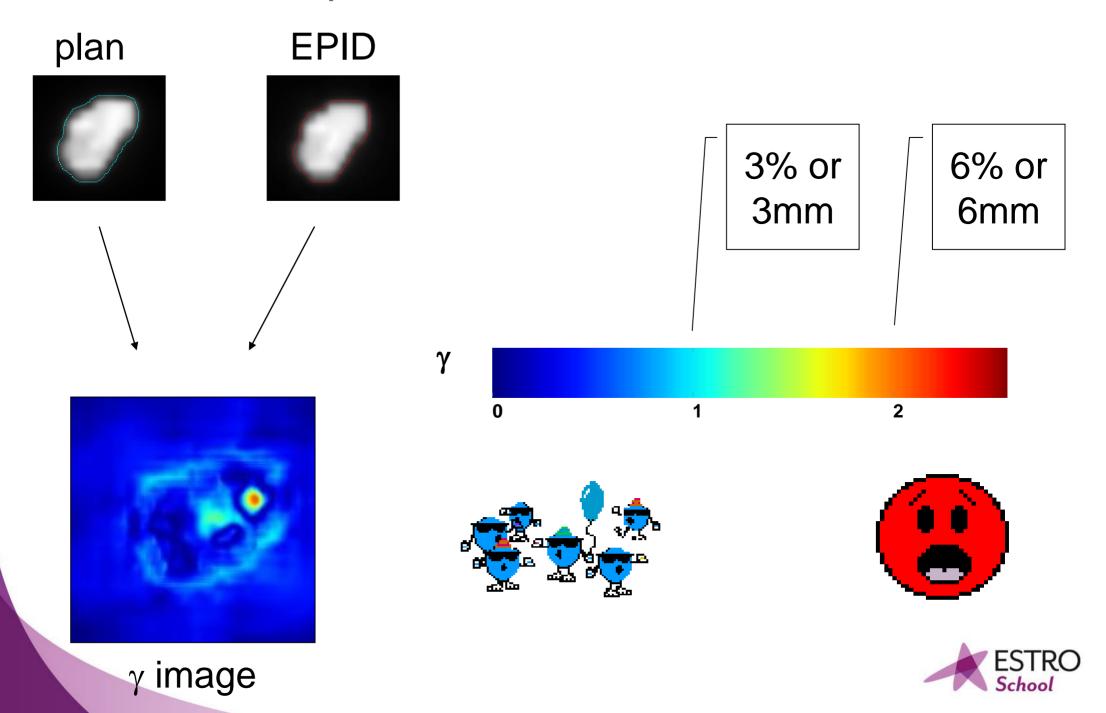


γ-evaluation: calculation vs measurement





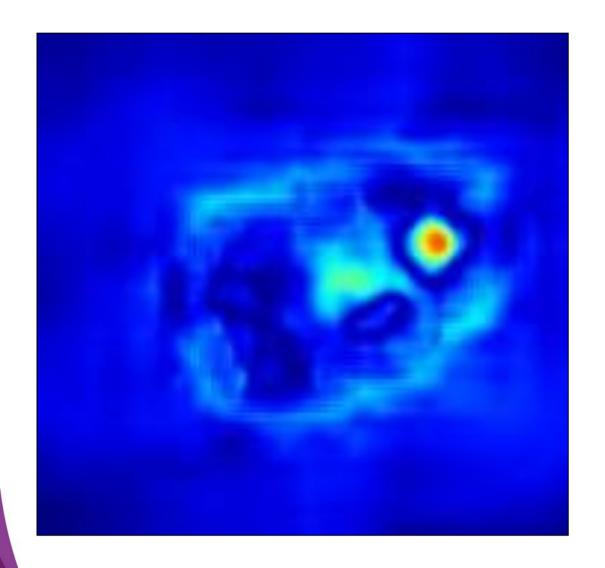
To compare the dose in 2D

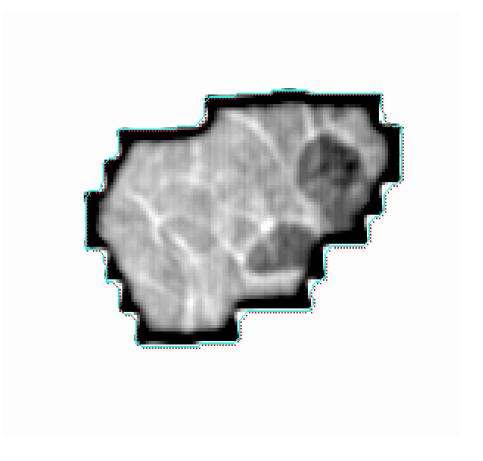


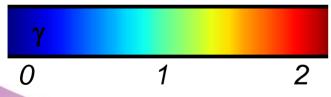
What can you detect?



Gas pockets

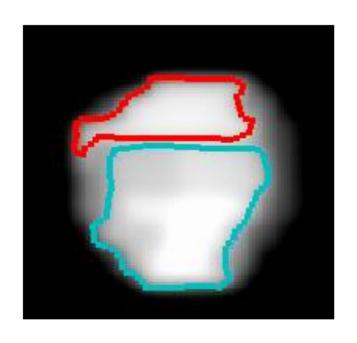




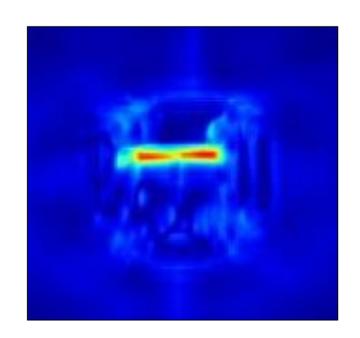




abutting leaves



isodose lines segments 3 & 6



γ-evaluation 3% / 3mm EPID vs plan



More errors?

- Transfer errors (planning → linac)
- Linac errors (both dosimetric and geometric)
- Dosimetric errors in plan
- Patient setup (e.g. CT reference to isoc shifts)
- Input errors
- Select the right patient / treatment in all systems



Patient setup

- CT reference to isocenter shift
 - Potentially really large errors (e.g. 10cm!)
 - They DO occur
- Possible countermeasures
 - Online imaging for ALL patients
 - Table shift surveillance software



ZERO



LCS: B2 PATID:

Please align patient

to CT Ref

Automatically retrieved from planning system



Includes shifts from offline protocols as well

Height:

a al

Longitudinal

-10.0

2.4

-3.1

TABLE:

Height:

Lateral:

Longitudinal:

0.0

0.0

0.0

ZERO



Interlock released when numbers are the same

Height:

Lateral:

Longitudinal

-10.0

2.4

-3.1

TABLE:

Height:

Lateral:

Longitudinal

-10.0

2.4

-3.1

Input errors / patient / treatment selection

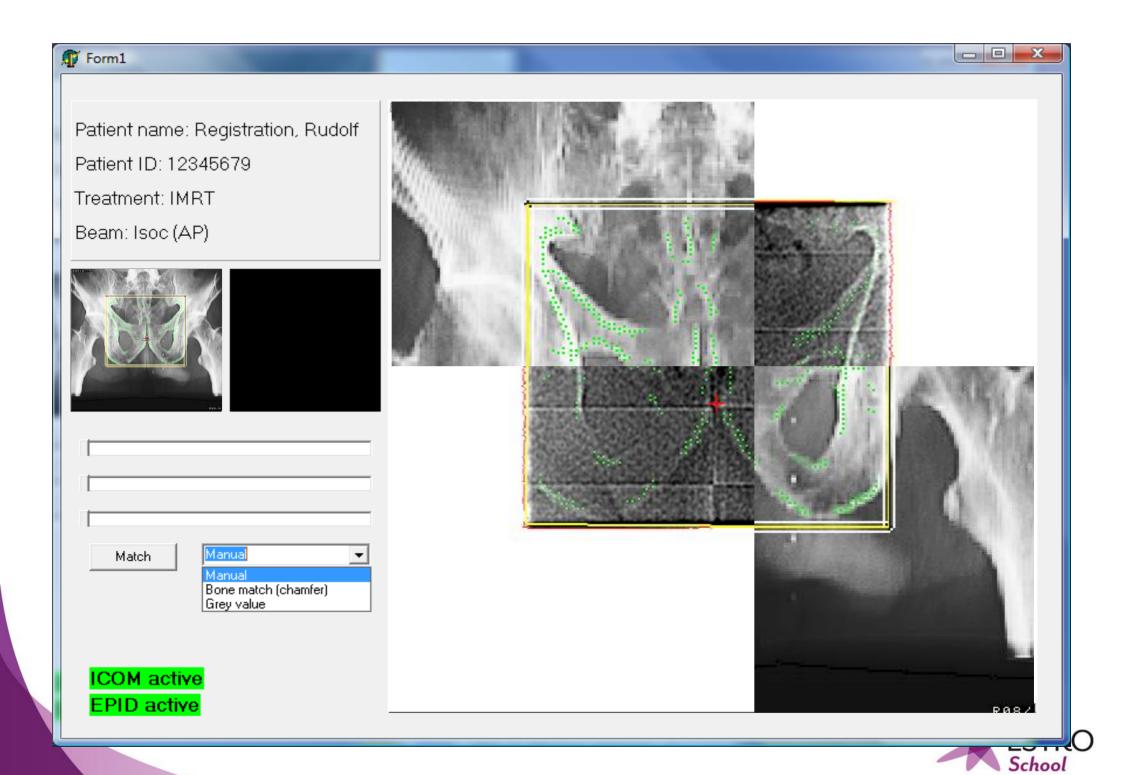
- Automation. Make the number of user interaction as small as possible
- Intuitive user interfaces
- Double checks
- New technology, like RFIDs?



Automation: EPID acquisition

- Radiographer...
 - Deploys the imager
- Application...
 - Selects patient and beam
 - Saves data in database without any user intervention
- Different screens, depending on beam property, e.g.
 - Dosimetry screen
 - Online registration screen
 - Breathhold screen





- - X X BREATHHOLD CHECK Patient name: van Vliet Patient ID: 12345678 Treatment: Breast breathhold Beam: Left lateral



Automation: Zero button EPID dosimetry

- Radiographer...
 - Deploys the imager and treats the patient
- Application...
 - 'Triggers' on new images from EPID acquisition application
 - Computes dose
 - Sends a report to physics
 - Notifies physics when something is wrong



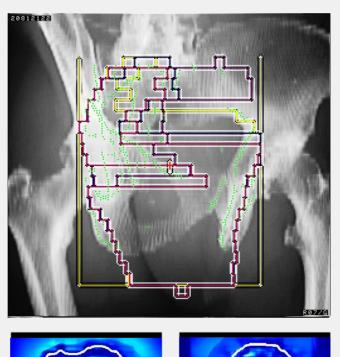


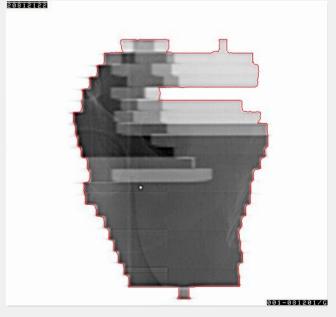
Patient name: Dosimetry, Dwayne

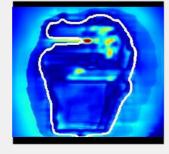
Patient ID: 12345679

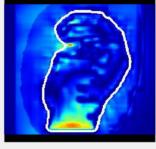
Treatment: IMRT

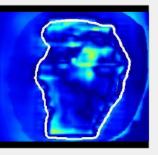
Beam Complicated one (7 of 7)

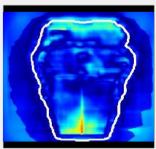




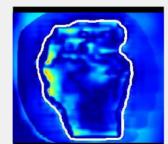


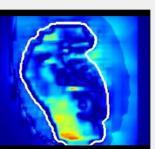


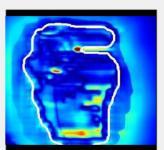




- 0 X







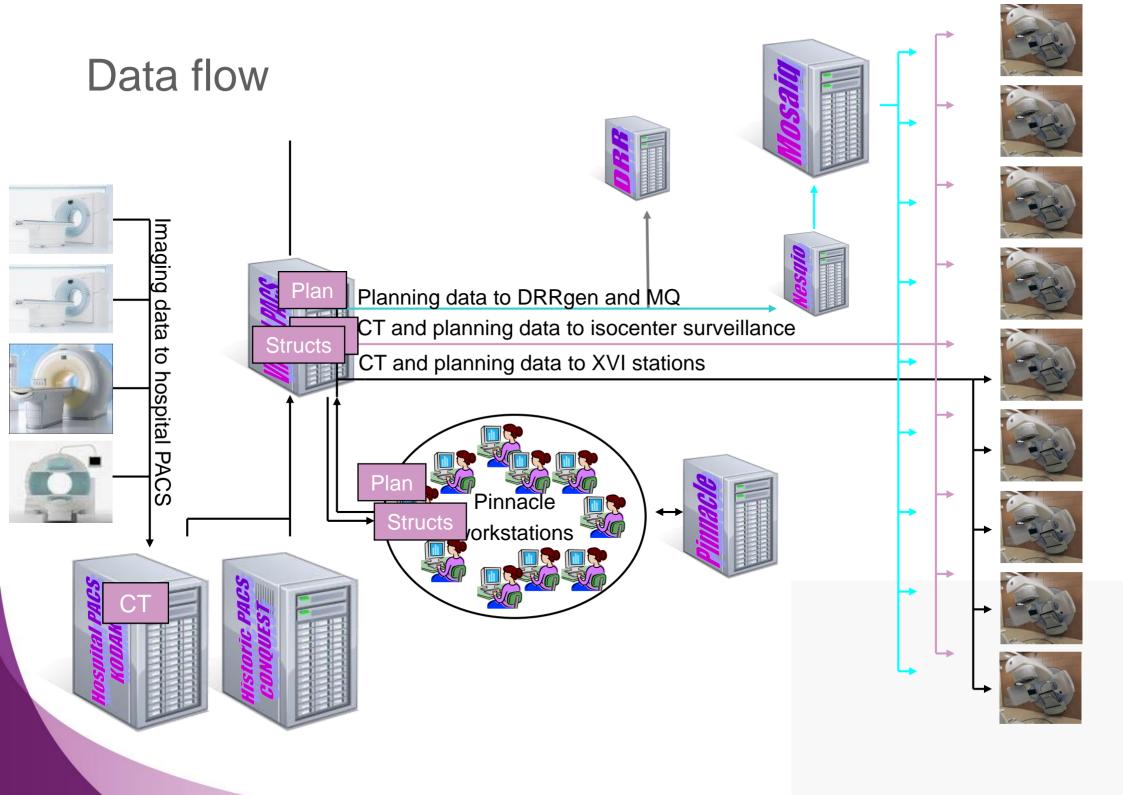


ICOM active EPID active

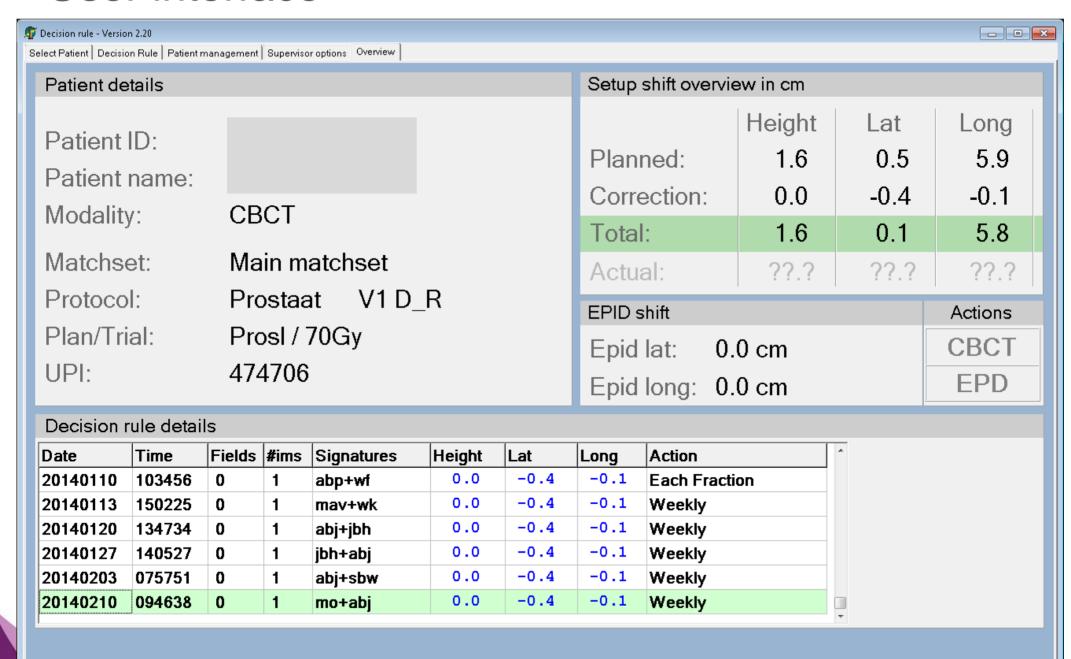
Automated dataflow example

- Dosimetrist sends plan for linac B5 to central server
- Server finds corresponding CT scan and structure set
- All data is then automatically sent to XVI station on B5
- Plan is sent to Mosaiq
- Plan and structures are sent to hospital PACS
- DRRs are automatically generated
- Patient is automatically entered in imaging database





User interface



Take home messages

- IGRT is good but not enough
- Take countermeasures to catch gross errors
- Try to find the simplest workflow (user interface, protocols, forms)
- Be especially aware when introducing new systems, protocols, or technologies





Incident management





Mirjana Josipovic

Dept. of Oncology, Rigshospitalet & Niels Bohr Institute, University of Copenhagen Denmark

Advanced skills in modern radiotherapy
June 2017



Intended learning outcomes

- Define an incident in radiotherapy context
- Discuss the importance of an incident reporting system
- Analyse the potential causes for an incident to have happened



Incident

• Any unintended event, including operating errors, equipment failures, initiating events, accident precursors, near misses or other mishaps, or unauthorized act, malicious or non-malicious, the consequences or potential consequences of which are not negligible from the point of view of protection or safety.

(IAEA Safety Glossary, 2007)

Radiation incident

• The delivery of radiation during a course of RT is other than intended by prescription, and could have or did result in unnecessary harm to the patient.

(Towards safer radiotherapy, BJR 2008)

Incident

• An *unplanned*, *undesired* event that hinders completion of a task and may cause injury, illness, or property damage or some combination of all three in varying degrees from minor to catastrophic. Unplanned and undesired do not mean *unable to prevent*.

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Incident

Any unintended event, including operating errors, equipment failures, initiating events, accident precursors, near misses or other mishaps, or unauthorized act, malicious or non-malicious, the consequences or potential consequences of which are not negligible from the point of view of protection or safety. Unintended Safety Glossary, 2007)

Radiation incident does not mean

The delivery of radiation during a course of RT is other than intended by prescription unable atorprevent! necessary harm to the patient.

(Towards safer radiotherapy, BJR 2008)

Incident

An unplanned, *undesired* event that hinders completion of a task and may cause injury, illness, or property damage or some combination of all three in varying degrees from minor to catastrophic. Unplanned and undesired do not mean unable to prevent.

Incidents

Actual incident = accident:

• The unforeseen event, that has affected the treatment of the patient

Potential incident:

- "Near miss"
- The unforeseen event, that was discovered and halted before it affected the treatment of the patient



From IAEA database of radiation incidents

Independent calculation checks 1998-2003 on 27830 charts/plans

An unintended "potential incident" was found:

- in ~3 % of all plans, during primary check
- in $\sim 1/2$ % of all plans, during secondary check

Actual incidents = accidents:

• in ~1/4 % of cases



For each actual incident, ~14 potential incidents were found through checking.

An incident frequency of 3% could be seen in a "typical clinic".

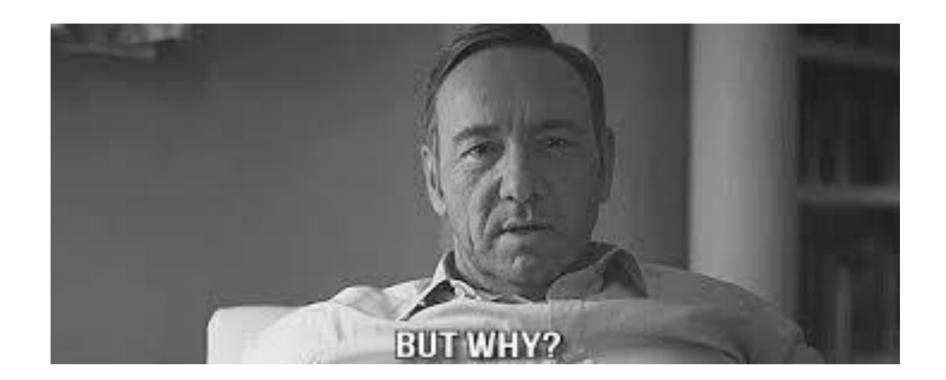


Incidents are more numerous than accidents:

• there are more opportunities to learn and improve the safety, than by only looking at major accidents.



But we do have a check procedure...

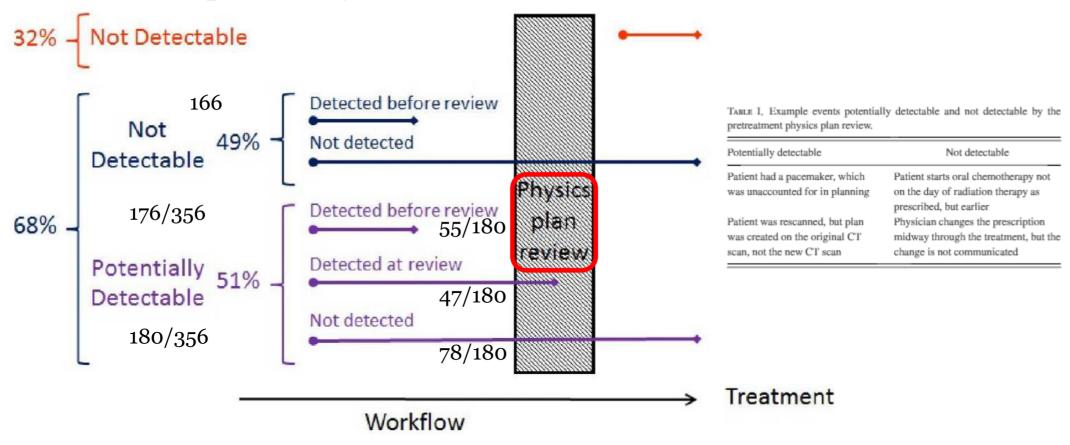




Incident frequency in modern radiotherapy

3011 reported incidents from 2012-2015 in single institution

552 potentially severe or critical

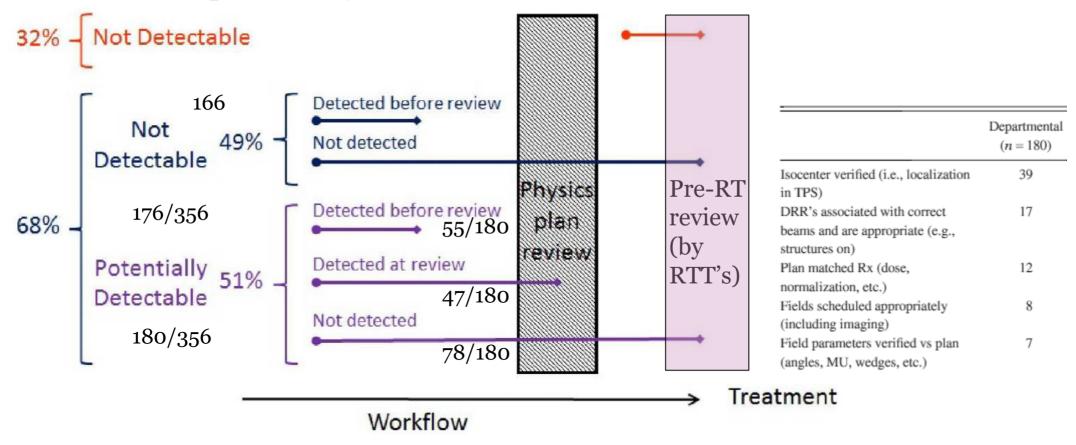




Incident frequency in modern radiotherapy

3011 reported incidents from 2012-2015 in single institution

552 potentially severe or critical



• Majority of potentially severe incidents occure before physics review $(68\%) - \sim 1/3$ of them is detected by review



Incident frequency in modern radiotherapy

3011 reported incidents from 2012-2015 in single institution

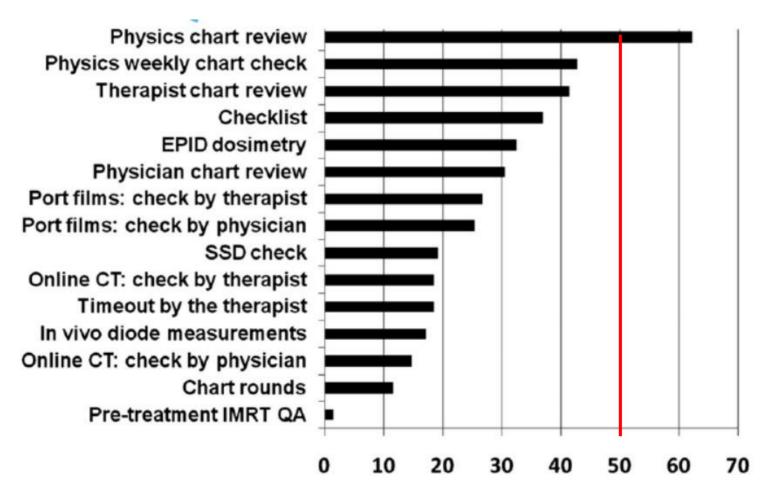
TABLE III. The percentage of potentially detectable and all events from the institutional ILS, which originated and were found at each step in the radiation therapy process.

Workflow step	% of potentially detectable events originating at this step	% of potentially detectable events found at this step	% of ALL events originating at this step	% of ALL events found at this step
Patient assessment	7.7	0.6	22.4	3.5
Simulation	28.2	3.3	13.0	8.2
Treatment planning	49.2	26.5	29.6	18.9
Plan review	1.7	38.1	4.7	22.3
Treatment delivery	2.8	14.9	8.9	29.1
On-treatment QM	1.6	8.8	2.8	9.4
Post-tx completion	0	0.6	11.4	6.6
Equipment and software QM	2.2	6.1	0.3	1.4
Not-defined	6.6	1.1	6.8	0.5

- Manual checks
- Majority detected by plan review need for improvement
- Recommendation for automatisation of check procedures



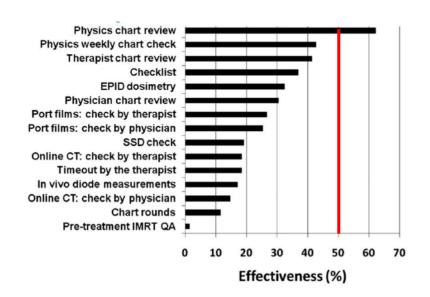
Are the check tools / procedures effective?



Effectiveness of a SINGLE check procedure [%]



Are the check tools / procedures effective?

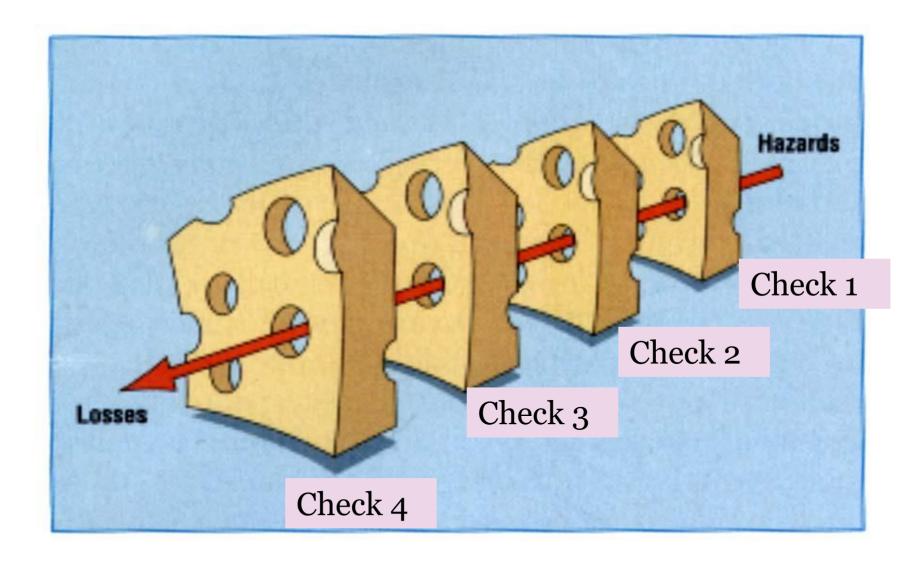


Combined effect of check procedures:

• 7 checks → 97% effectiveness



Swiss cheese model of accident causation





Many incidents have a variable magnitude:

- same type of incident can have different impact on different patients / treatment sites
- next time the same incident happens, it may become an accident



Incident prevention to improve patient safety

Proactive

- Patient safety rounds
- Leadership tool

Reactive

Reporting and analysing incidents



- Blaming individuals is emotionally more satisfying than targeting institutions
- We cannot change the human condition, but we can change the conditions under which the humans work



• Incident reporting must not result in disciplinary investigation as a consequence of reporting





internal



- locally
- inside your dept / institution

external

- outside your organisation
- sharing with peers

mandatory

to regulatory authorities



voluntary

 to professional (inter)national organisation



internal

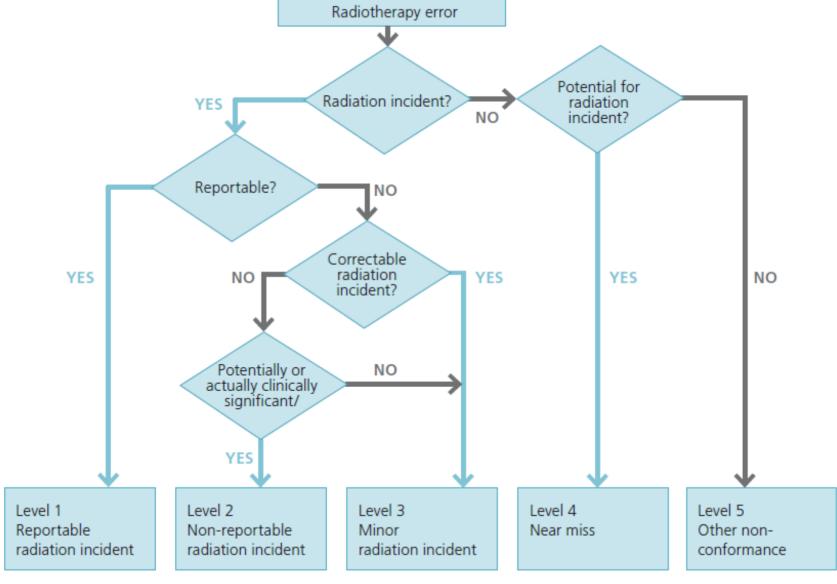


external

- Bigger "pool of events" facilitate better identification of safety critical steps in the process of radiotherapy
- Incidents from another hospital can lead to early identification of hazard in your own hospital, before an actual incident occurrence
- Providing general culture of safety awareness



What to report?





What to report?

You should report all unintended incidents:

- Observed by you, during involvement in the incident
- Observed by observing others
- Made to attention at a later point in time

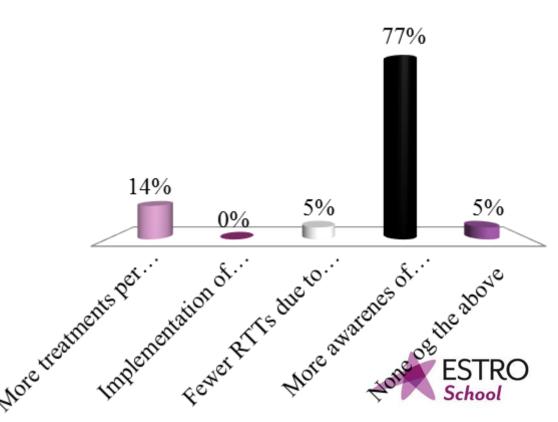
All incidents
affecting patient safety
or
potentially affecting patient safety



How to explain the increase of incidents?

Year	2008	2009	2010	2011
# of incidents reported	14	30	115	122

- A. More treatments per linac
- B. Implementation of advanced technology
- C. Fewer RTTs due to budget cut downs
- D. More awarenes of incident reporting
- E. None og the above



Role of incident reporting system

- To **identify** system design flaws and critical steps in the radiotherapy pathway
- To highlight critical problems and patterns of causes of these problems
- To spread knowledge on new risks or involving new technology
- To **promote safety culture** and awareness through involvement of and feedback to staff and managers
- To prevent repeated incidents



Role of incident reporting system

Incident reporting system has to be a part of a longer chain:

- Incident Identification
- Reporting
- Investigation
- Analysis
- Management
- Learning



Analysis methods

- Root cause analysis
- Journalaudit
- Mortality analysis
- Global Trigger Tool





Root cause analysis

A systematic method to identify

- WHAT happened ...the actual chain of events leading to the incident
- WHY could it happen ...identification of what caused the incident
- HOW to prevent the incident to happen again ...action plan & follow up

•...NEVER, who caused the incident



Take home message

- Incidents are more numerous and varying than actual accidents
- By learning from the incidents happening in your clinic you can avoid a potential future accident
- Incident report is an essential tool for safer radiotherapy







Workshop on Plan Selection

Rianne de Jong *RTT*Academic Medical Centre, Amsterdam
Prague 2017



Contents workshop

Plan selection strategies (IGRT-part); theoretical background and AMC experience

Bladder:

- Registration protocol for bladder
- Live Observer Study

Cervix:

- Registration protocol for cervix
- Live Observer Study

Rectum:

- Registration protocol for cervix
- Live Observer Study



Flavours in selection structures

- GTV/CTV
- PTV
- 95% (dose lines)
- ITV
- PTV + 3mm



Plan selection at AMC

Bladder

- Start 2013
- Selection based on CTV
- Full and empty bladder CT

Cervix

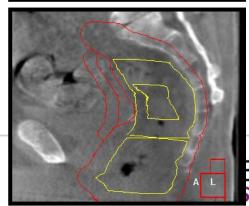
- Start 2015
- Selection based on ITV
- Full and empty bladder CT

Rectum

- Start 2016 March
- Selection based on variable margins
- Full bladder CT only







Plan selection at AMC

Bladder

- Start 2013
- Selection based on CTV
- Full and empty bladder CT

Cervix

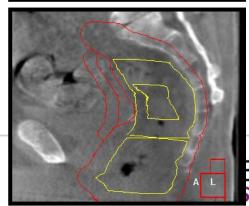
- Start 2015
- Selection based on ITV
- Full and empty bladder CT

Rectum

- Start 2016 March
- Selection based on variable margins
- Full bladder CT only







Plan selection Rectum

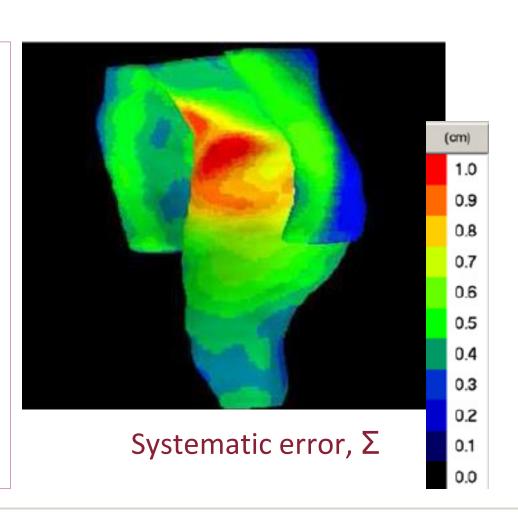
Largest uncertainty:

Upper-anterior side

No correlation with bladder but <u>rectum filling!</u>

Choice & Number margins:

- Encompass largest uncertainty
- Feasible workload for treatment planning
- Complexity of selection at Linac



Implementation Strategy

Initial strategy plan selection: straightforward

- Only 1 CT scan no extreme bladder/rectum fillings
- No need of algorithm

But personally, concerns:

- More complex target definition target volume compared to bladder and cervix?
- Larger patient numbers more RTTs to be trained
- Dosimetric benefit given anatomy of target and OAR

Implementation strategy:

- ✓ Observer study to assess feasibility and support training and implementation (large number RTTs) *
- ✓ Dosimetric study to assess dose to OAR **



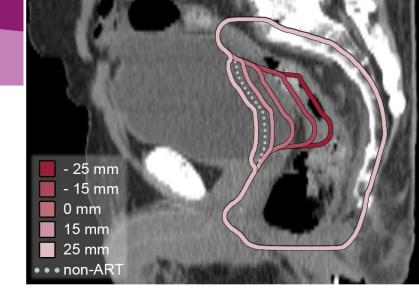
Design of the observer study

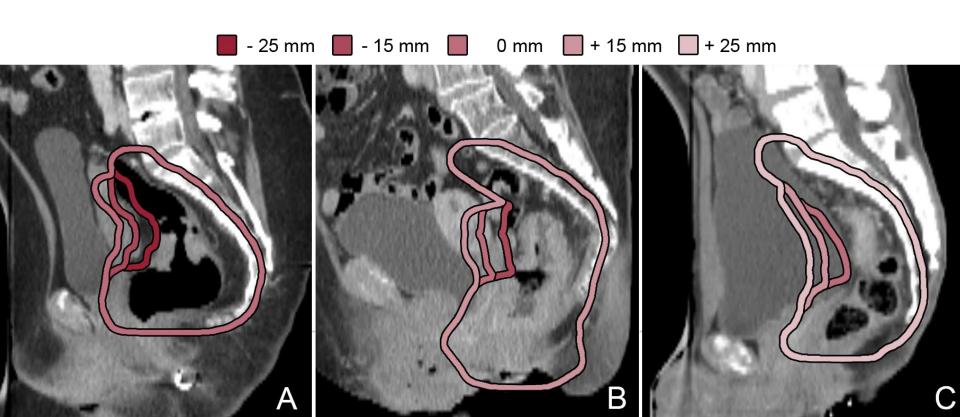
- Select the tightest fitting plan -
- 20 observers
 - 15 RTT's, 1 PhD student, 4 Physicists
- 11 patients
 - 5 prone with BB, 6 supine
- 5 CBCT scans per patient
- Delineations according to Roels et al, 2006, IJROPB
- 3 margins for ventral side upper mesorectum depending on anatomy captured on CTref:

Rectum	Full	Medium	Empty
Margins	-25 mm, -15 mm, 0 mm	-15 mm, 0 mm, 15 mm	0 mm, 15 mm, 25 mm

Margins plan selection

Visual representation of the variable margins to ventral upper side mesorectum:





Design of the study

1. Lecture on target definition by expert Radiation Oncologist



2. First measurement - single observers



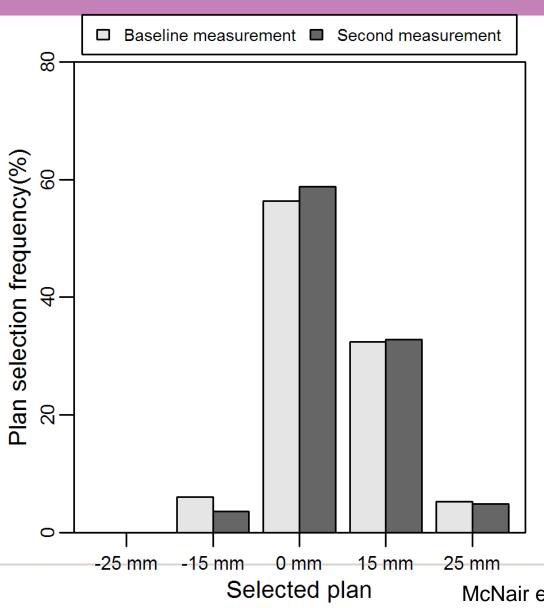
3. Consensus meeting - reviewing all patients, all CBCT scans with all observers and expert radiation oncologists: defining the gold standard



4. Second measurement - single observers



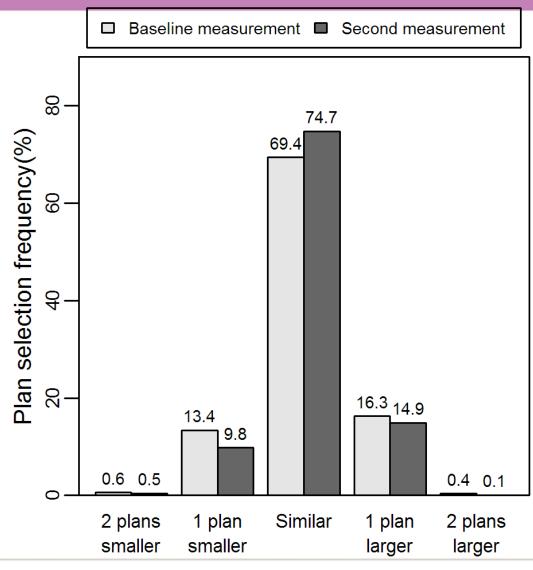
Outcome of the study

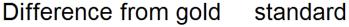


Non adaptive margin 20mm

McNair et al. (2015) Br J Radiol. EST

Outcome of the study







Outcome of the study

Did we select a too small plan fewer times 2nd time around?

- Yes,
$$15,6\% - 10.3\%$$
 ($p = 0.002$)

Did we select a too big plan fewer times 2nd time around?

- No,
$$16.6\%$$
 - 15.0% ($p = 0.130$)

Was there a difference between supine and prone BB?

- Maybe, in favor of supine measurement 1 70.0% versus 64.2% p > 0.05 measurement 2 74.2% versus 67.0% p > 0.05



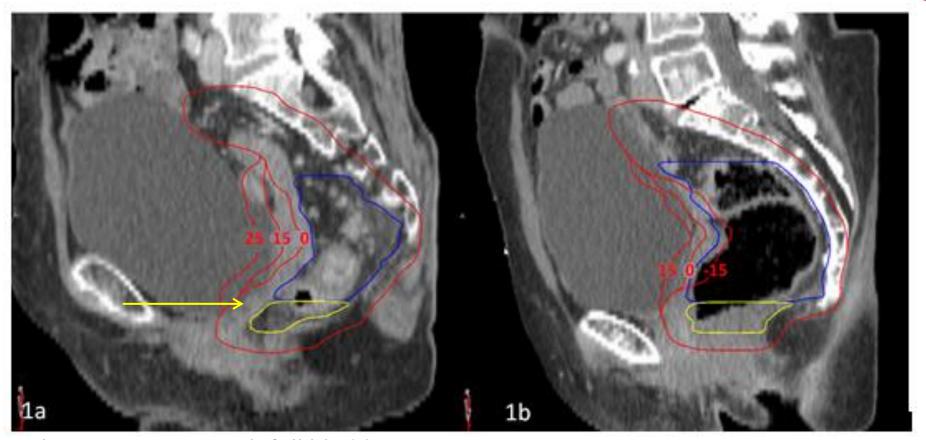
Implementation - conclusion

Observer study:

- Plan selection is feasible: 75% concordance with gold standard
- Observer study also served as training tool
- Margin of -25 mm never selected



Clinical Protocol plan selection Rectum



1 Planning CT scan with full bladder

A. Empty rectum on planning CT: 25 mm, 15 mm, 0 mm anterior margins

B. Full rectum on planning CT: 15 mm, 0 mm, -15 mm anterior margins



Clinical protocol plan selection Rectum

- 2 sets of 3 margins
- Long (25x2Gy) and short (5x5Gy) treatment
- VMAT
- Daily CBCT
- 1/w post treatment CBCT: intra fraction motion
- 1/w retrospective review: all plan selections

consistency imaging- and management system



Clinical protocol plan selection Rectum

Exclusion criteria:

- Dual hip prostetics
- T4 tumors
- Time constraints can not be met
 - Inguinal lymfe nodes
 - Length CTV_Mesorectum_up < 5 cm in CC direction



Plan selection at the treatment machine:

First week: 1 trained RTT and 1 physicist, 1 physician

Second week: 2 RTTs (1 trained)



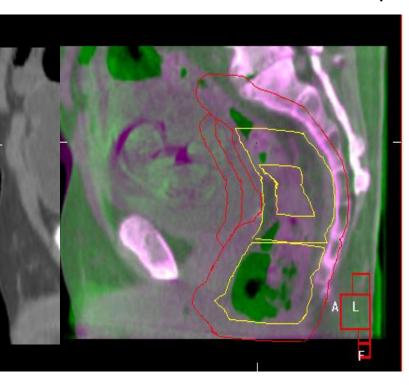
Target volume on Planning CT



Plan selection at the treatment machine:

First week: 1 trained* RTT and 1 physicist, 1 physician

Second week: 2 RTTs (1 trained)



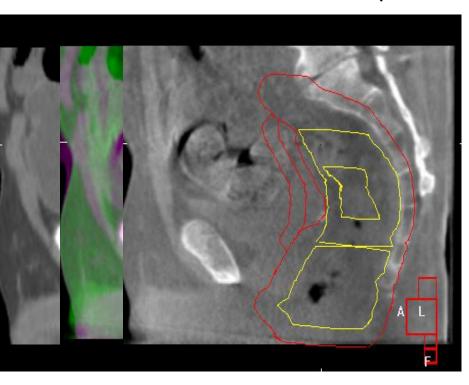
Bone match
Overlay CT/CBCT

^{*} de Jong et al.(2016), Radiother. Oncol.

Plan selection at the treatment machine:

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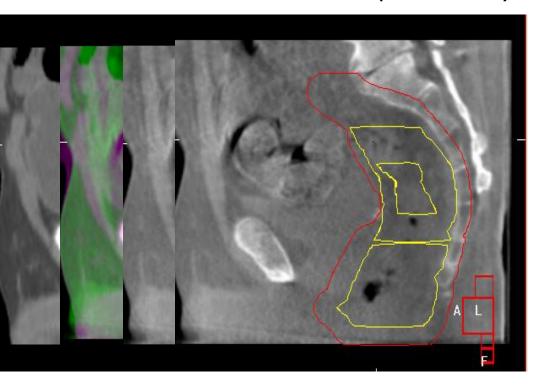
Target volume & margins on CBCT

^{*} de Jong et al.(2016), Radiother. Oncol.

Plan selection at the treatment machine:

First week: 1 trained* RTT and 1 physicist, 1 physician

Second week: 2 RTTs (1 trained)



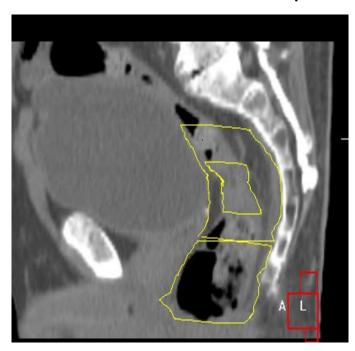
Selected margin
On CBCT

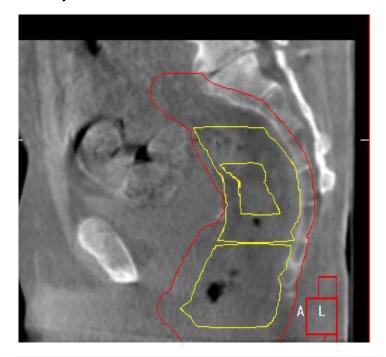
^{*} de Jong et al.(2016), Radiother. Oncol.

Plan selection at the treatment machine:

First week: 1 trained* RTT and 1 physicist, 1 physician

Second week: 2 RTTs (1 trained)





Empty bladder but smallest margin!



March 2016 – May 2017

70 patients treated with plan selection

Evaluation of the first 20 (consecutive) patients

10x short treatment scheme (5x5Gy)

10x long treatment scheme (25x2Gy)

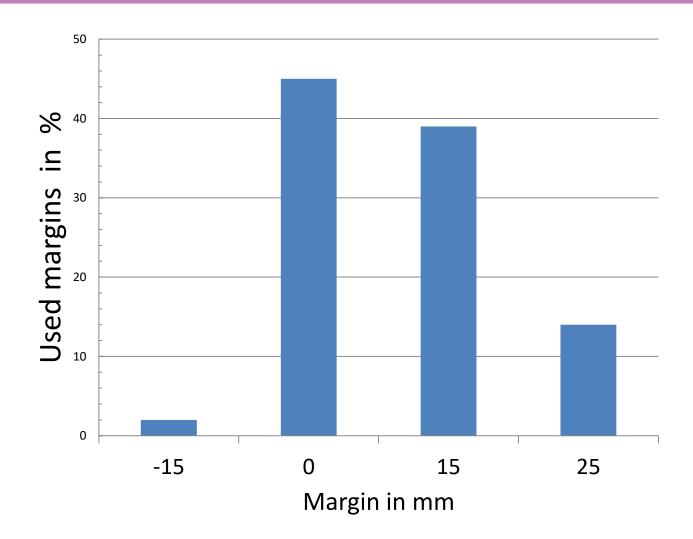
Margins sets used:

Full rectum (+15 / 0 / -15mm) 30%

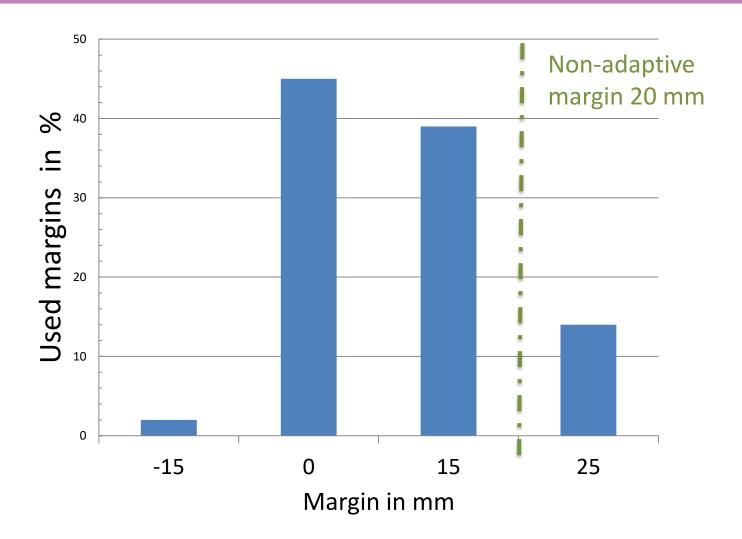
Empty rectum (+25/ +15 / 0 mm) 65%

Full rectum (+15 / 0 mm) 5% (insufficient TP time)

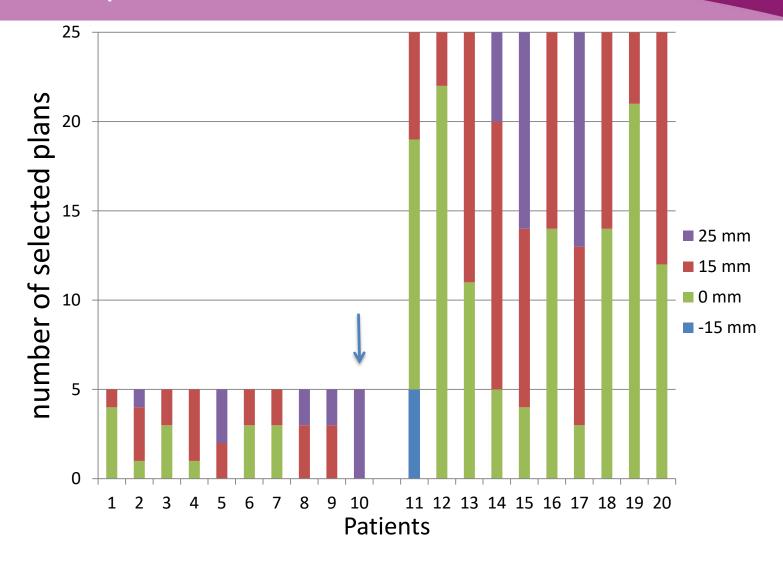




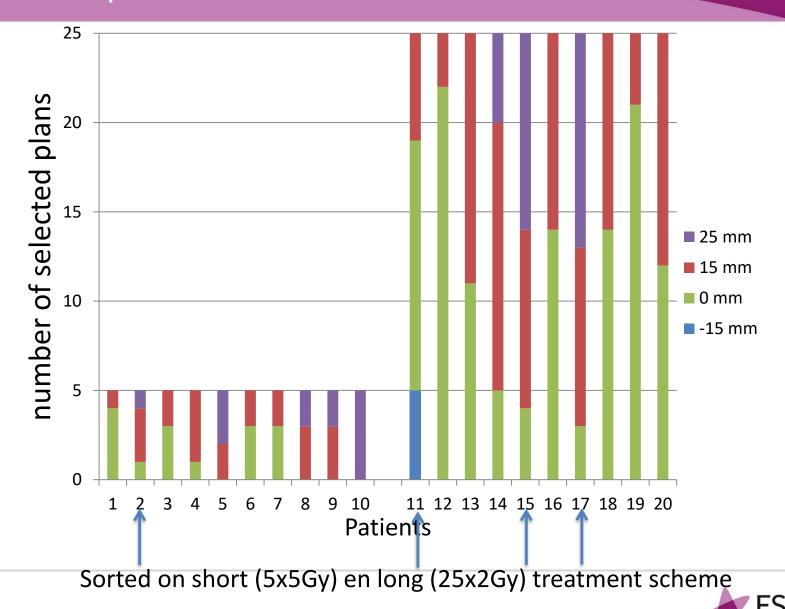






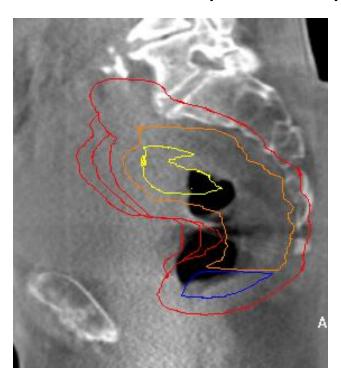


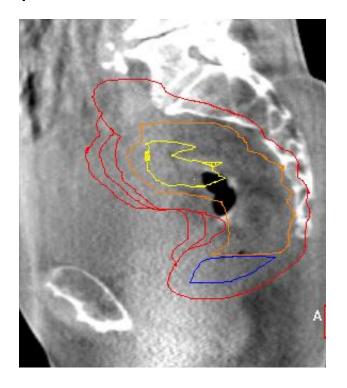
Sorted on short (5x5Gy) en long (25x2Gy) treatment scheme



Delayed treatment: $7 \times (5 \times in 1 \text{ patient})$

To obtain a more favorable anatomy in case of a very full rectum, usually caused by gas pockets







Delayed treatment: 7 x (5 x in 1 patient)

To obtain a more favorable anatomy in case of a very full rectum, usually caused by gas pockets

Post-treatment CBCT 1pw:

1 fraction the selected plan was no longer suitable due to a moving gas pocket



Delayed treatment: $7 \times (5 \times in 1 \text{ patient})$

To obtain a more favorable anatomy in case of a very full rectum, usually caused by gas pockets

Post-treatment CBCT 1pw:

1 fraction the selected plan was no longer suitable due to a moving gas pocket

The weekly review:

Smaller margin could have been selected in 20% of fractions, and a larger margin in 2% of fractions

No inconsistencies between the imaging system and radiotherapy management system!



Summary evaluation

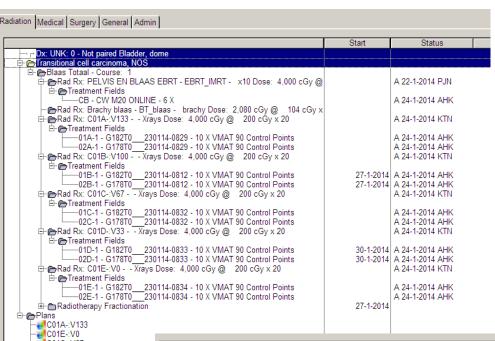
Plan selection for rectum cancer with variable margins for upper mesorectum for first 20 patients:

- Both sets of margins used
- Majority of patients needed multiple margins
- Limited influence of intra fraction motion
- Good consistency in weekly review
- No errors between imaging and management system
- Limited delay remains due to anatomy on CBCT

- Successfully and safely implemented! -



Management system - Mosaiq



Not optimized for library of plans

- 1. Empty sessions
- 2. Sessions with all plans available

- € C01C-:V67	Session Setup / Field									Notes Sts By 1:Rx:C01A-:V133				1:Rx:C	1B-:V100		1:Rx:C01C-:V67 1:			:Rx:C01l				
	No	Date	Time	ID	Tx ED	Seq		Meterset	Dos	e Machine	TSPFDC		-,	Fx E		Cum	Fx ED	Dly	Cum	Fx ED	Dly (Cum F	x ED	Ī
└ <mark>.</mark> C01D-:V33	⊕-QA 2	7-01-2014								AMC-U3	S												\Box	_
	₽-1			2Flds			1PI				TS						1	200 cGy	200 cGy					
		8-01-2014					1PI				TS						2 1	200 cGy	400 cGy					
		9-01-2014					1PI			AMC-U3	S						3 2	200 cGy	600 cGy					
		0-01-2014					1PI 1PI			AMC US	S												1	
		3-02-2014 3-02-2014					1PI			AMC-U3 AMC-U3	0													
		4-02-2014					1PI			AMC-U3	S													
		5-02-2014					1Pi			AMC-U3	Š													
		6-02-2014					1PI			AMC-U3	S													
	₫- 10	7-02-2014	9:00	0Flds			1PI			AMC-U3	S													
		0-02-2014					1PI			AMC-U3	S													
		1-02-2014					1PI			AMC-U3	S													
		2-02-2014					1PI			AMC-U3	S													
		3-02-2014 4-02-2014					1PI 1PI			AMC-U3 AMC-U3	5													
		7-02-2014					1PI			AMC-U3	S													
		8-02-2014					1Pi			AMC-U3	S													
		9-02-2014					1PI			AMC-U3	S													
	d- 19 2	0-02-2014	9:00	0Flds			1PI			AMC-U3	S													
	<u>⊞</u> -20 2	1-02-2014	9:00	0Flds			1PI			AMC-U3	S													



Extra checks at AMC

ART - Rectum

Patiënt naam:	
Patiënt nummer:	2650976
Course:	1
Plannen:	-15 tot +15

Gegevens opslaan

						Post CB	CT	1				
Fractie	Laborant(en)	Datum	Plan	Х	Υ	Z		Opmerkingen	Week	Eval.	Door	Opmerkingen
1 2 3 4 5	mms Iba ahk mka ahk jwa ahk nwn ahk/pen/awn eac/awn/wls	30-05-16 31-05-16 01-06-16 02-06-16 03-06-16	0 +15 0 0 +15	-0,02	-0,02	-0,08	Y	geen fysicus bereikbaar in Amsterdam! zaten te twijfelen tussen 0 en +15	1			
6 7 8 9 10									2			
11 12 13 14 15									3			
16 17 18 19 20									4			
21 22 23 24 25									5			
D 0 (-1 004)	C -::			DT	7			- II- K	IOADT			

Per 8 februari 2016 zijn we begonnen met planselectie voor rectumpatiënten: ART rectum. Zie voor complete werkinstructie Kwadraet 'Rectum ART'. ledere dag moet er iemand van de IGART-groep aanwezig zijn bij planselectie.

NB: In tegenstelling tot ART-Blaas mag er bij ART-Rectum NOOIT getweakt worden!

Vragen en/of opmerkingen horen wij graag. Alvast dank.

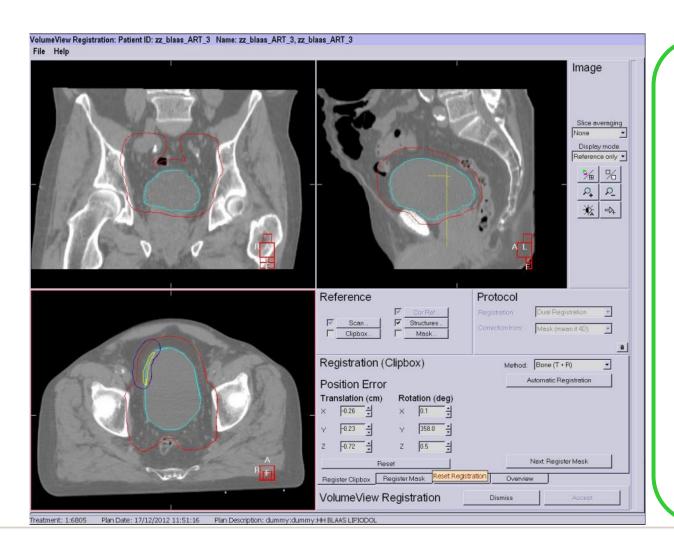
Jorrit (62594) Rianne (66857)



Live plan selection

Bladder





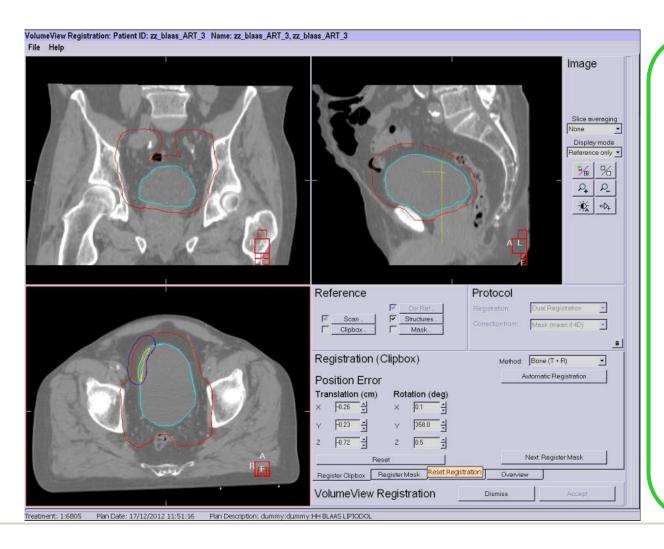
Bladder plan selection @AMC

Targetvolume:

- Whole bladder low dose
- Boost part bladder high dose
- Nodal area up to L5

VMAT delivery

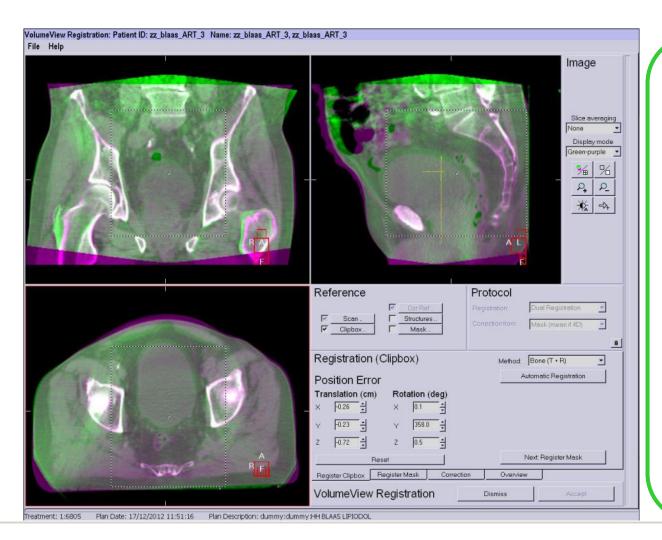




Bladder plan selection @AMC

- Full bladder protocol
- Bony anatomy registration for nodes
- 3. Selection of plan for whole bladder
- 4. Optional: tweak for the high dose region





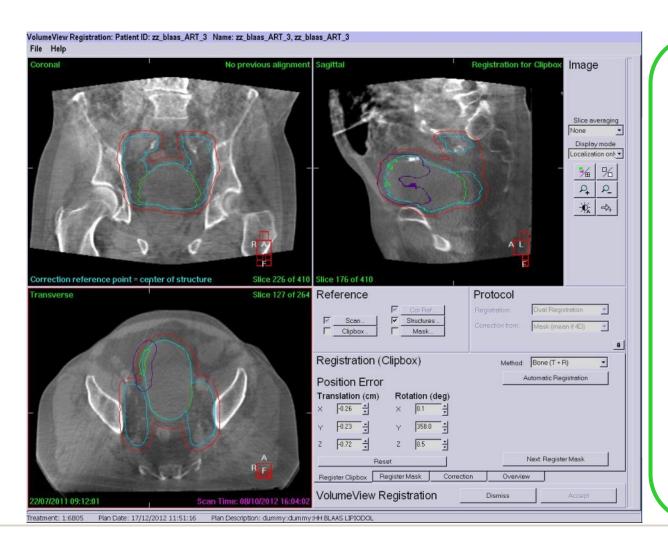
 Bony anatomy registration in green-purple with bone algorithme





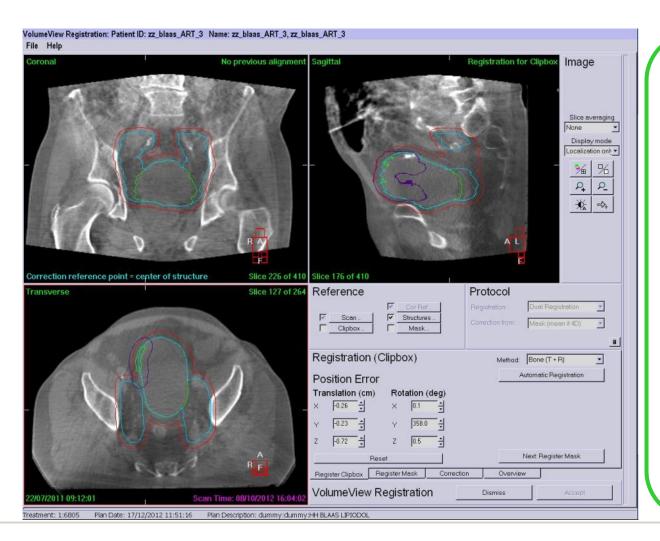
- Overlay of structures on the CBCT
- Do not use CT scan anymore at this point!
- Pick the plans that fits best based on bladder structure, by means of deduction





- Display accompanying structures
- Check target coverage in PTV, both before and after correction for rotations





Optional tweak: manual adjustement for the high dose region



Do not overstep on the tweak. Take the marginsize of the nodes into account



Live Observer Study

4 bladder patients x 3 CBCT's

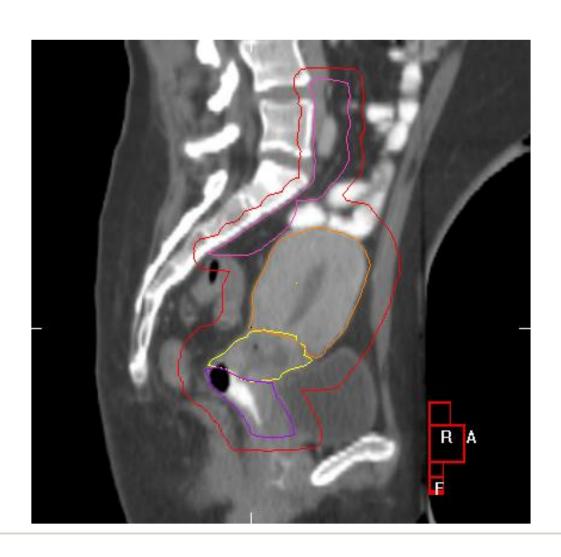
- 1. Individual selection by turning point
- 2. Group discussion
- Selection by turning point



Live plan selection

Cervix





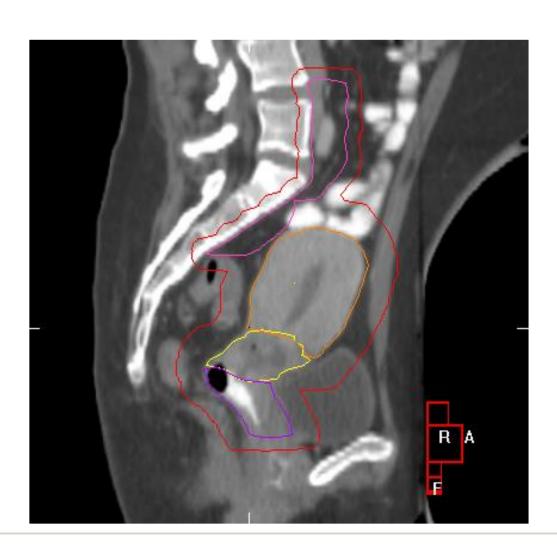
Cervix plan selection @AMC

Target volume:

- Cervix
- Uterus
- Nodal region up to L2

VMAT delivery

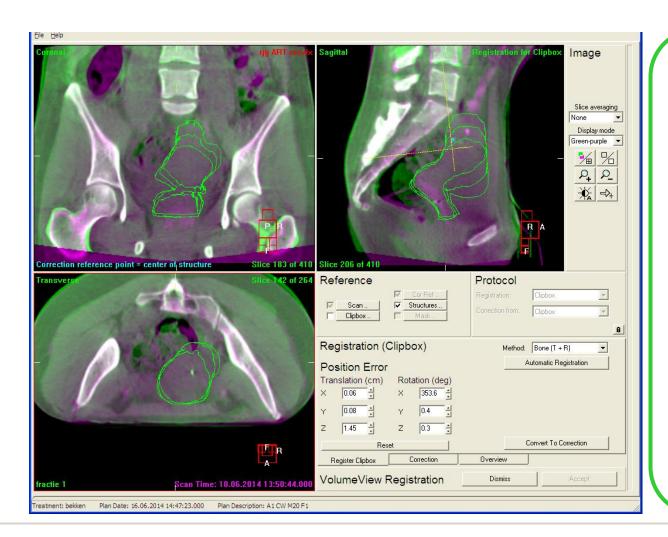




Cervix plan selection @AMC

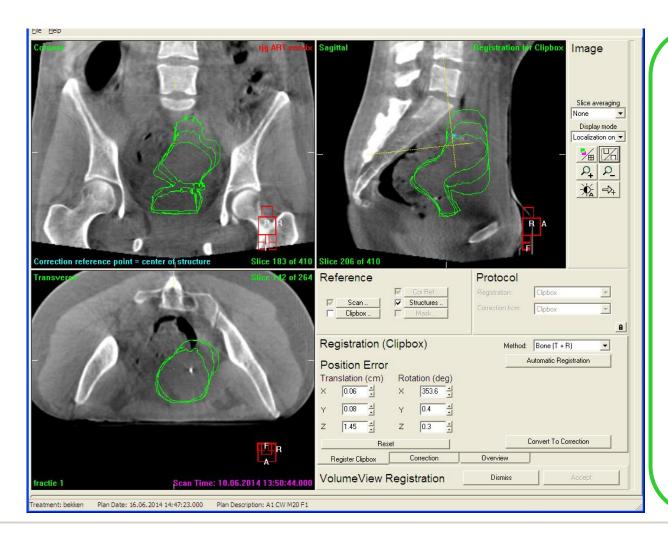
- Full bladder protocol
- 2. Bony anatomy registration for nodes
- 3. Selection of plan for cervix&uterus
- 4. Marker check
- 5. NO tweak





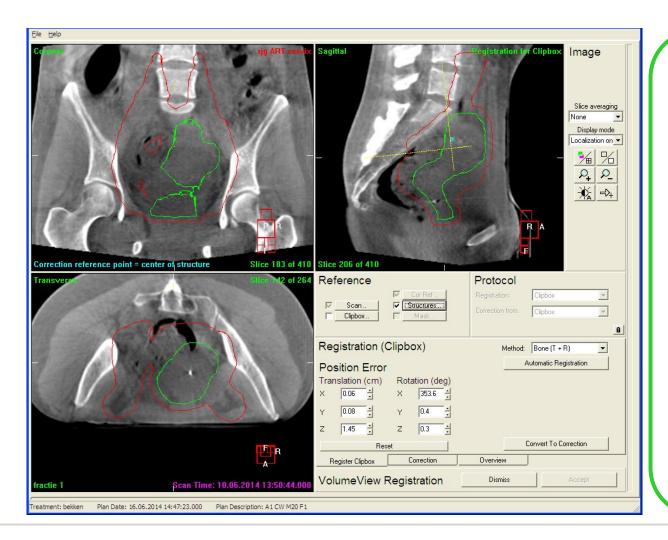
Bony anatomy registration in green purple overlay with bone algorithm





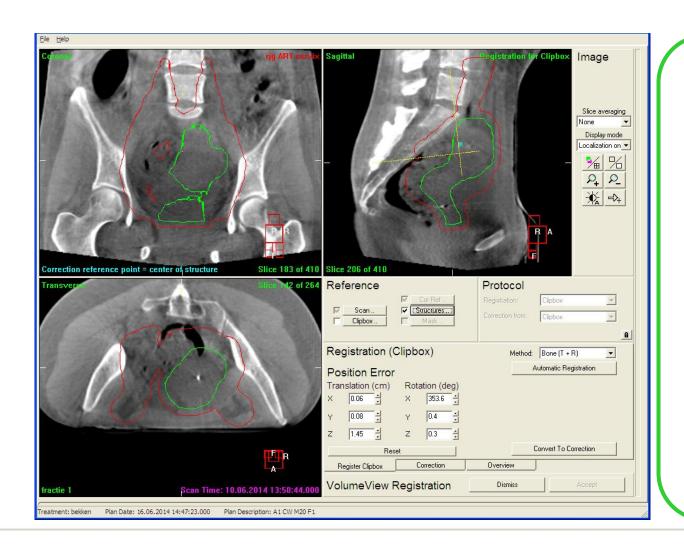
- Overlay of (ITV) structures on the CBCT
- Do not use CT scan anymore at this point!
- Pick the plans that fits best based on ITV structure, by means of deduction





- Display accompanying structures
- Check target coverage in PTV, both before and after correction for rotations
- Check markers





No Tweak!!



Live Observer Study

4 cervix

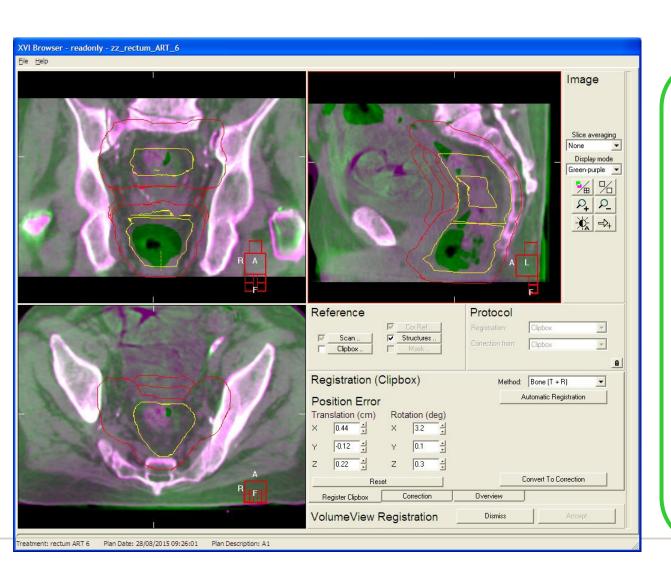
- 1. Individual selection by turning point
- 2. Group discussion
- Selection by turning point



Live plan selection

Rectum





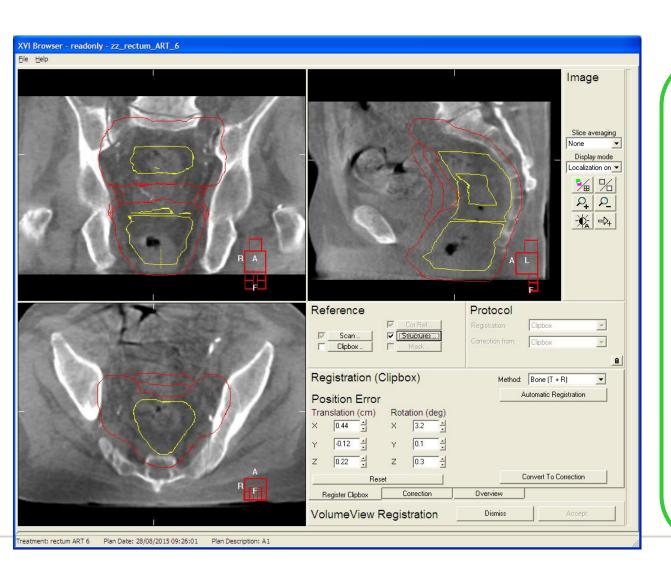
Bony anatomy registration (T+R)





Inspection of delineation of upper mesorectum and lower mesorectum at reference CT scan





Select thigthest fitting plan (margin) based on CBCT





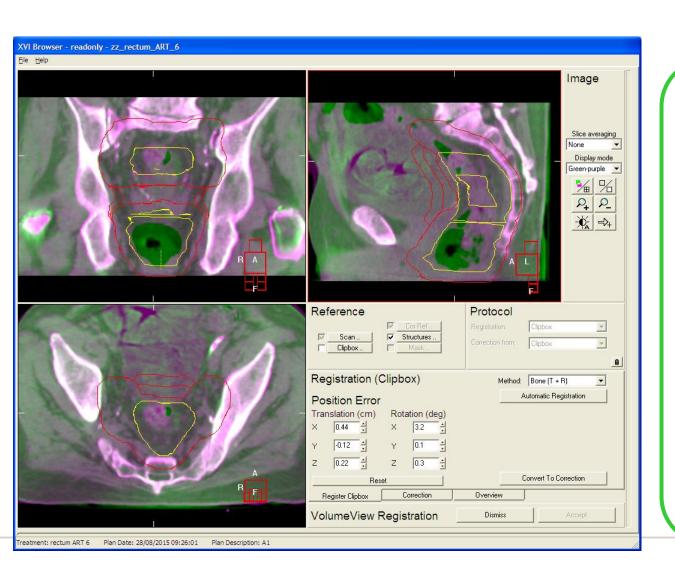
Check coverage again after recalculation of rotations

NO TWEAK!

Think lymfe nodes







1

Bony anatomy registration (T+R)

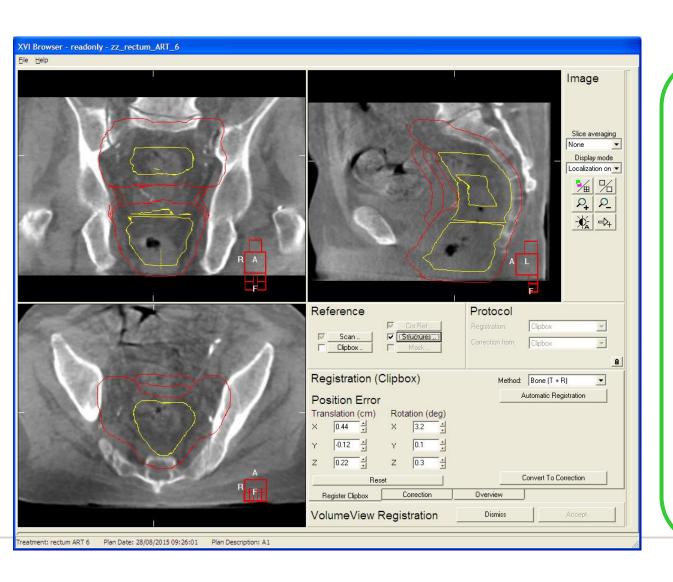




2

Inspection of delineation of upper mesorectum and lower mesorectum at reference CT scan

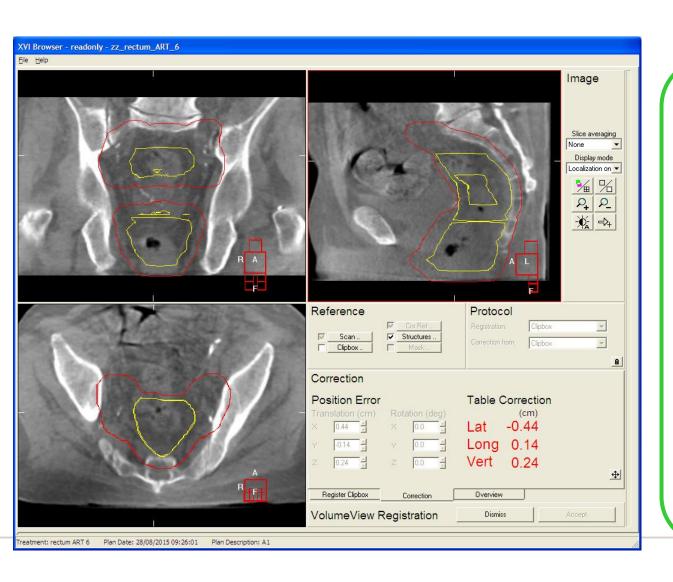




3

Select thigthest fitting plan (margin) based on CBCT





4

Check coverage again after recalculation of rotations



Prospective Adverse Event Reporting and the Role of the RTT

Liz Forde, RTT
Assistant Professor
The Discipline of Radiation Therapy
School of Medicine
Trinity College Dublin







Toxicity in Oncology

- Toxicity and tolerance differs for each organ
- Toxicity has an undeniable impact on patients psychosocial well being and quality of life
- Factors impacting on toxicity and patient tolerance:
 - Biological
 - Subjective
 - Duration of reaction
 - Response to medical intervention



Toxicity in Oncology

Acute reactions

- During or shortly after treatment
- Common for epithelial tissue damage
- > Typically temporary
- > Support through the most severe phase
 - Medical intervention
- Psychosocial
- Do not ignore unexpected acute toxicities

Late reactions

- Months or even years following treatment
- > Too late for a change in treatment
- Often in deeply seated organs
- Clinical observation difficult







Need for Recording and Reporting

- Survival and success stories frequently reported
- Adverse events and poor outcome data rarely reported
- Large variation in grading, analysing and reporting
 - Standardisation is required
 - Comparison between trials, patients groups, institutions
- More combined therapies
- More aggressive therapies
- More complex treatment regimes

Associated with higher acute toxicity



Need for Recording and Reporting

- Routine reporting involves commitment to prospective documentation, analysis and long term follow up
- Culture of the department and education of staff
 - Radiation oncology vs. Medical oncology vs. Surgical oncology
 - Single modality vs. multi modality trials



The Four Domains of Adverse Event Reporting

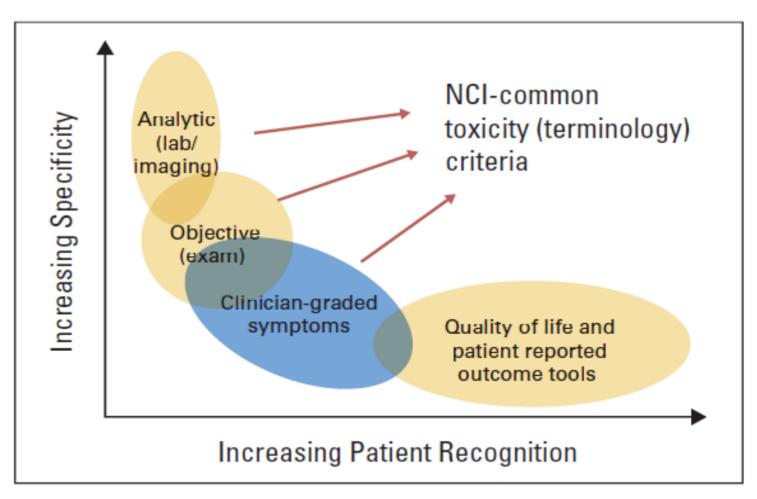


Fig 1. Adverse effects domains. NCI, National Cancer Institute. Adapted with permission.8

Assessment and Reporting of Adverse Events

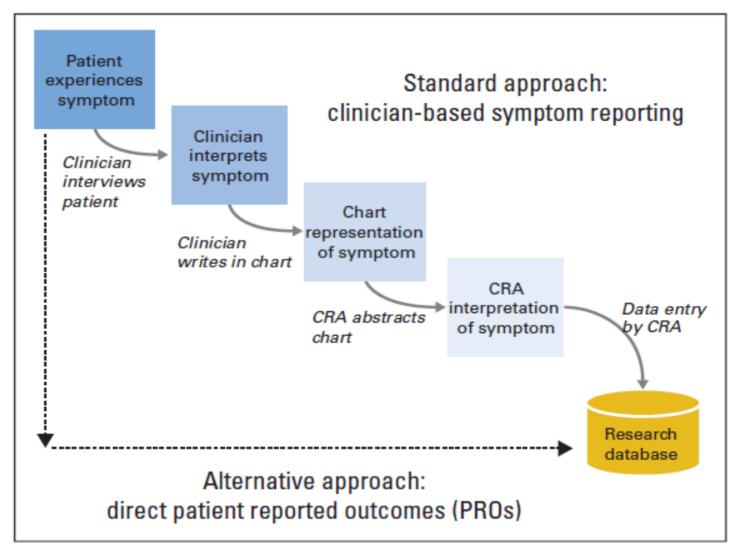


Fig 2. Flow of symptom information in cancer treatment trials. CRA, clinical research assistant. Reprinted with permission.³⁶



Features of a Scoring System

- Specific descriptions
- Unambiguous language
- Consistency and reliability
 - Decreased inter-user variation and misinterpretation
- Grading of severity
 - > Intervention required
 - > Impact on QoL or daily activities



Systems Developed

- WHO 1979
- NCI CTC 1983
- RTOG 1984
- RTOG/EORTC 1984
- Franco-Italian Dictionary
- The Dische grading dictionary
- LENT-SOMA 1995
- CTCAE Version 3 2003
- CTCAE Version 4 2010
- All with varying degrees of content and severity of scaling
- Need for standardisation and amalgamation of acute and late effects... EST

Chemotherapy only

Radiation Oncology, Acute Only

LENT SOMA

- Perception of toxicity between patient and physician can be very different
 - Irreversible
 - Protracted
 - Uncontrollable
 - Social debilitating
- Combination of data from functional tests and also a subjective score



The Work of the NCI

- CTC v1.0 developed in 1983
 - > Chemotherapy only
 - Acute reactions only

- CTC v2. updated in 1997
 - > Intended for *all oncology modalities*
 - > >250 descriptive criteria
 - > Still only addressed grading of *acute* toxicity



NCI - CTCAE v3.0

- 2003
- All organ systems covered with a total of 370 criteria listed
- Amalgamation of acute and late effects
- Can be applied to *all modalities* (Surgical, medical and radiation oncology)
- Duration and sequence of an adverse event should be recorded
- This is a "grading dictionary" not intended to assess treatment regimes or determine what is acceptable or not
 - This is still a clinical judgement of risks vs. benefits



CTCAE v4

- 2010
- Harmonise terminology with MedDRA
- Organisation of document changes
 - Version 3 was divided into categories based on either pathophysiology or anatomy
 - Version 4 is based on system organ class (SOC)
- Result: Decreased number of terms (1059 down to 790)
- FAQ document available online



Common Terminology Criteria for Adverse Events v4.0 (CTCAE)

Publish Date: May 28, 2009

Quick Reference

The NCI Common Terminology Criteria for Adverse Events is a descriptive terminology which can be utilized for Adverse Event (AE) reporting. A grading (severity) scale is provided for each AE term.

Components and Organization

SOC

System Organ Class, the highest level of the MedDRA hierarchy, is identified by anatomical or physiological system, etiology, or purpose (e.g., SOC Investigations for laboratory test results). CTCAE terms are grouped by MedDRA Primary SOCs. Within each SOC, AEs are listed and accompanied by descriptions of severity (Grade).

CTCAE Terms

An Adverse Event (AE) is any unfavorable and unintended sign (including an abnormal laboratory finding), symptom, or disease temporally associated with the use of a medical treatment or procedure that may or may <u>not</u> be considered related to the medical treatment or procedure. An AE is a term that is a unique representation of a specific event used for medical documentation and scientific analyses. Each CTCAE v4.0 term is a MedDRA LLT (Lowest Level Term).

Definitions

A brief definition is provided to clarify the meaning of each AE term.

Grades

Grade refers to the severity of the AE. The CTCAE displays Grades 1 through 5 with unique clinical descriptions of severity for each AE based on this general guideline:

- Grade 1 Mild; asymptomatic or mild symptoms; clinical or diagnostic observations only; intervention not indicated
- Grade 2 Moderate; minimal, local or noninvasive intervention indicated; limiting age-appropriate instrumental ADL*.
- Grade 3 Severe or medically significant but not immediately life-threatening; hospitalization or prolongation of hospitalization indicated; disabling; limiting self care ADL**.
- Grade 4 Life-threatening consequences; urgent intervention indicated.
- Grade 5 Death related to AE.

A Semi-colon indicates 'or' within the description of the grade.

Not all Grades are appropriate for all AEs. Therefore, some AEs are listed with fewer than five options for Grade selection.

Grade 5

Grade 5 (Death) is not appropriate for some AEs and therefore is not an option.

Activities of Daily Living (ADL)

- *Instrumental ADL refer to preparing meals, shopping for groceries or clothes, using the telephone, managing money, etc.
- **Self care ADL refer to bathing, dressing and undressing, feeding self, using the toilet, taking medications, and not bedridden.



CTCAE v4

• Example of AEs potentially experienced by prostate radiotherapy patients

	Grade 1	Grade 2	Grade 3	Grade 4	rade
Diarrhea	Increase of <4 stools per day over baseline; mild increase in ostomy output compared to baseline	Increase of 4 - 6 stools per day over baseline; moderate increase in ostomy output compared to baseline	Increase of >=7 stools per day over baseline; incontinence; hospitalization indicated; severe increase in ostomy output compared to baseline; limiting self care ADL	Life-threatening consequences; urgent intervention indicated	Death
Definition: A disorder characteriz	ed by frequent and watery bowel r	movements.	•		
Proctitis	Rectal discomfort, intervention not indicated	Symptoms (e.g., rectal discomfort, passing blood or mucus); medical intervention indicated; limiting instrumental ADL	Severe symptoms; fecal urgency or stool incontinence; limiting self care ADL	Life-threatening consequences; urgent intervention indicated	Death
Definition: A disorder characteriz	ed by inflammation of the rectum.				
Fatigue	Fatigue relieved by rest	Fatigue not relieved by rest;	Fatigue not relieved by rest,	-	-

Definition: A disorder characterized by a state of generalized weakness with a pronounced inability to summon sufficient energy to accomplish daily activities.



Even with advances in toxicity reporting using CTCAE variability still remains



Patient Reported Outcomes (PRO)

- HCP generally *underestimate* side effect presentation, severity and duration compared with patients
- Agreement is generally closer for observable side effects than for subjective ones
 - E.g. diarrhoea is observable and fatigue is subjective
- PROs cover the subjective domain
 - E.g. Pain
- Issues re literacy
 - Questionnaires to guide a consult is not considered a true PRO as there is still some level of interpretation and collection by someone other than the patient

PROs

- The NCI have since developed a web based PRO for the CTCAE
- 81 symptoms have been identified for inclusion in a PRO
- 126 questions assess the different attributes of these symptoms
- Language has been adjusted for patients
 - Myalgia is "translated" as aching muscles

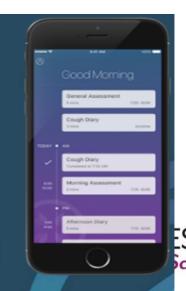


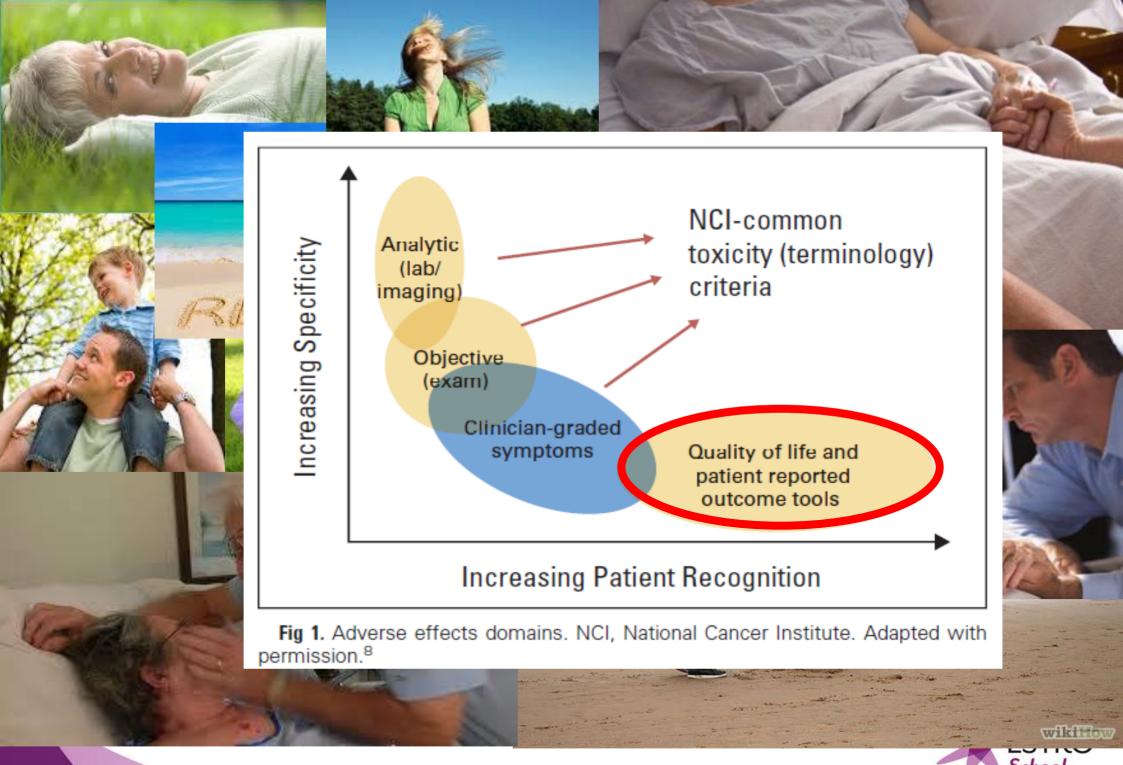
ePRO

- Basche presented at ASCO in June 2017
 - > >700 patients treated at MSKCC
 - Breast and lung
- "Real time" reporting of side effects
- Web based PRO for chemo patients
- Works on smart phones
- Nurses get sent an email when side effects worsen

Maximise patient reported outcomes with ePRO

Empower your patients & boost Patient Reported Outcomes with ePRO from IBM Clinical Development, Watson Health.





"To the clinician and the biologist the preservation of functions that are essential to life would seem of paramount importance. But to the patient, the obligation to live a long and painful existence may be worse than death itself. The economic consequence of being unable to work, and even more, being utterly dependent on others for day to day activities like feeding, dressing and washing are not easy for a third person to appreciate. Similarly facial disfigurement and anal or bladder incontinence may impose such social consequences on the patient that may become effectively housebound even though their other vital organs function, motor activities and pain threshold are virtually unimpaired."



Quality of Life Assessment

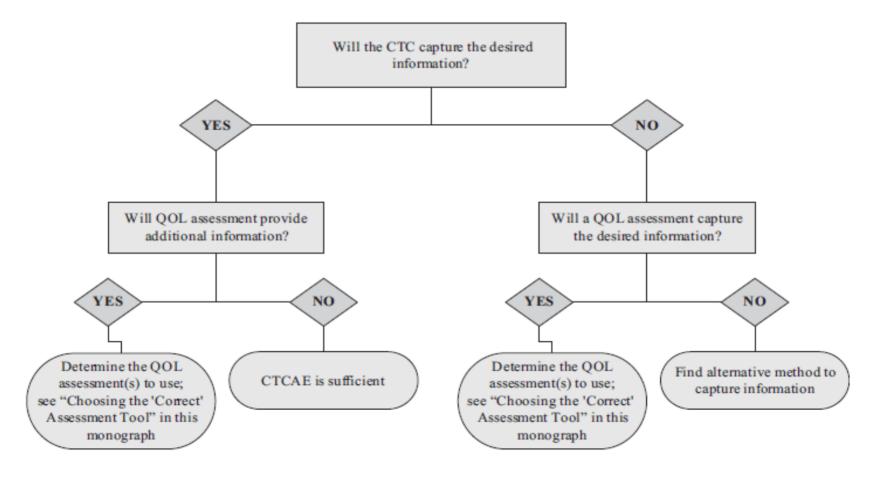


FIG 2. Flow diagram to determine the method for adverse event and QOL data collection.

Huschka M and Burger K. Does QOL provide the same information as toxicity data? Curr Probl Cancer. 2006; 30(6): 244-254



QoL Assessment

- QoL is *subjective* and depends on patients ability to adapt to a certain extent
 - QoL measures not the AE itself but the how it impacts on daily activities
- QoL includes psychosocial support networks and patient's spirituality
- QoL and AE reporting are complimentary to each other
- This combination strengthens the patient physician relationship
 - Recognition of different goals
 - Overall survival, but at what cost



QoL Assessment

- QoL assessment also lacks consistency between trials, countries, departments and patient groups
- Assessment Scales available
 - > The Symptom Distress Scale
 - ➤ The Lung Cancer Symptom Scale
 - Functional Assessment of Chronic Illness Therapy Diarrhoea
 - > The International Prostate Symptom Score
 - 8 questions includes 1 QoL question
 - ➤ The Expanded Prostate Cancer Index Composite (EPIC)
 - Urinary
 - Bowel
 - Sexual function
 - Hormonal changes



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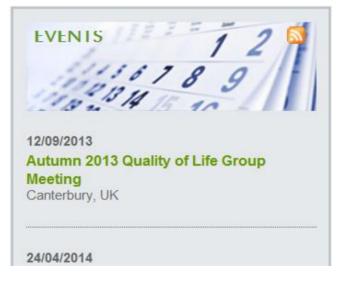
QUEST

The EQuestion to asset cancer

The EOR life of can



1 2 3 4 5





Careers & Grants

QoL Assessment

- QLQ C30
- Current version = version 3
- Translated into 81 languages
- 3000 studies internationally
- Disease specific modules also available for use:
 - Breast, Lung, Head & Neck, Oesophageal, Ovarian, Gastric, Cervical cancer, Multiple Myeloma, Oesophago-Gastric, Prostate, Colorectal Liver Metastases, Colorectal and Brain



During the past week:	Not at All	A Little	Quite a Bit	Vo M
17. Have you had diarrhea?	1	2	3	
18. Were you tired?	1	2	3	
19. Did pain interfere with your daily activities?	1	2	3	
20. Have you had difficulty in concentrating on things, like reading a newspaper or watching television?	1	2	3	
21. Did you feel tense?	1	2	3	,
22. Did you worry?	1	2	3	
23. Did you feel irritable?	1	2	3	
24. Did you feel depressed?	1	2	3	,
25. Have you had difficulty remembering things?	1	2	3	
26. Has your physical condition or medical treatment interfered with your <u>family</u> life?	1	2	3	
27. Has your physical condition or medical treatment interfered with your social activities?	1	2	3	4
28 Has your physical condition or medical treatment				

Some Limitations of Scoring Systems in General

- Inconsistencies in the timing of data recording
- Time consuming and resource intensive data collection
- Transfer of information and data collection
 - > Interpretation of information from patient to clinician
 - Manually entered into database
- Underreporting of lower grades (Grade 1 and Grade 2)



Is There a Role for the RTT?



Treatment Review Clinics

- Clinical examination
- Side effects are explained and assessed
- Medication or intervention may be required
- Nutritional advice
- CAM advice
- Psycho social issues are addressed
- Documentation of intervention and progress
- Unrelated medical advice
- Quality assurance for the progression of treatment
- Logistical information

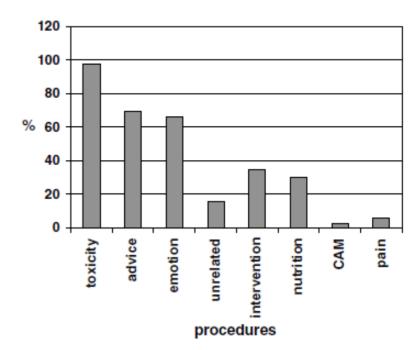


Fig. I. Frequency of procedures observed during treatment reviews. Toxicity, toxicity scoring indicated; Advice, advice on side effects; Emotion, emotional support given in terms of assurance and information given; Unrelated, unrelated medical problems addressed; Intervention, medical intervention given in the form of drug prescription, liaison with other physicians, ordering of wound dressing, or any other investigations; Nutrition, nutritional advice given; CAM, complementary and alternative medicine addressed; Pain, pain score taken.

Shi et al., 2009



Table I Medical intervention rates versus treatment site observed in Phase 2

Treatment site	Breast	Brain	H & N	Thorax	Pelvis	Other	Total	Pearson's Chi-square
Number	11	6	13	8	15	3	56	
(% within site)	(19%)	(54.5%)	(40.6%)	(38.1%)	(65.2%)	(20%)	(35%)	P = 0.001

Shi et al., 2009

Table 1. Breakdown of treatment review clinics requiring medical intervention (MI) and no MI according to the site of the cancer being treated.

Treatment site	MI required n (%)	No MI required n (%)	Total clinics n (%)
Head and neck	41 (93)	3 (7)	44 (22)
Prostate	11 (28)	29 (73)	40 (20)
Chest	18 (78)	5 (22)	23 (12)
Rectum	13 (59)	9 (41)	22 (11)
Breast	7 (33)	14 (67)	21 (11)
Brain	8 (73)	3 (27)	11 (6)
Gynaecological	10 (91)	1 (9)	11 (6)
Bladder	3 (33)	6 (67)	9 (5)
Superficial	2 (33)	4 (67)	6 (3)
Bone metastases	2 (40)	3 (60)	5 (3)
Pelvis	2 (50)	2 (50)	4 (2)
Abdomen	1 (50)	1 (50)	2 (1)
Extremity	0 (0)	2 (100)	2 (1)
Total clinics	118 (59)	82 (41)	200 (100)

As an initial step limit RT lead review to sites of low MI



Table III ROs' and RTTs' concerns with regards to RTT-led treatment reviews				
ROs' concerns	RTTs' concerns			
(1) Training [9]	(1) Medico-legal responsibility [21]			
(2) Scope of practice — RTTs must know when to refer to ROs [7]	(2) Training [18]			
(3) Medico-legal responsibility [5]	(3) Resource, time and manpower constraints [16]			
(4) Resource, time and manpower constraints [4]	(4) Remuneration [14]			
(5) Patients' perspective [2]	(5) Support from ROs and management [14]			
(6) Compromise in RTT work performance due to diversification of role [1]	(6) Patients' perspective [13]			
(7) Overconfidence of RTTs [1]	(7) Increase workload for RTTs [12]			
	(8) Lack of licensing — prescription, decision making, recognition for leading reviews [6]			
	(9) Sensitivities of job overlap with nurses and ROs [5]			

Shi et al., 2009

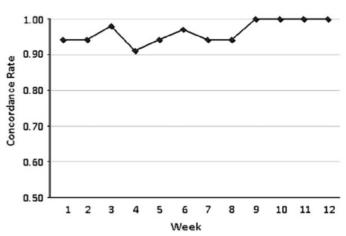


Figure 2. Average weekly concordance rates between the clinical specialist radiation therapist and radiation oncologist.

Lee et al., 2012



Impact of This Approach?

For the *Individual*?

- Increased job satisfaction
- Mutual respect as a professional
- Specialisation
- Autonomy in the workplace
- Personal growth
 - Career advancement in a field that has a historical "ceiling"



- Improved MDT dynamics
- Increased efficiency
- Better use of staff skills
- Education of peers
 - Mentorship





Take Home Messages

- Diligent adverse event reporting should not be reserved for clinical trials
- **Prospective** data collection that is electronic and easily accessible
- Language needs to be clear for *all* members of the team accessing patient notes
- Better equipped to assess impact of treatment in an evidence based approach

