

AAPM 2021 April 17-20 | VIRTUAL
 SPRING CLINICAL MEETING

Safety Considerations for Proton Therapy

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Promises of proton therapy

beam direction →

X-ray (photon) beam

proton beam #1

SOBP region (12 proton beams)

#2

#3

skin

shallow tissues

tumor

deep tissues

From Wikwand

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Sensitive to WET variation

10 MV Photon

Proton SOBP

1.0 cm WED Variation

3%

90%

Dose (arbitrary unit)

Water Equivalent Depth (WED)

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Potential reasons for WET variation

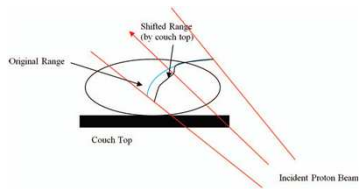
- ▶ Potential range perturbations caused by the immobilization device
- ▶ Setup error
- ▶ Inter and intra-fractional motion
- ▶ Patient anatomy change
 - ▶ Tumor regression
 - ▶ Patient weight loss
 - ▶ Development of edema
 - ▶ Bladder and bowel filling variation

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Range perturbations caused by the table top



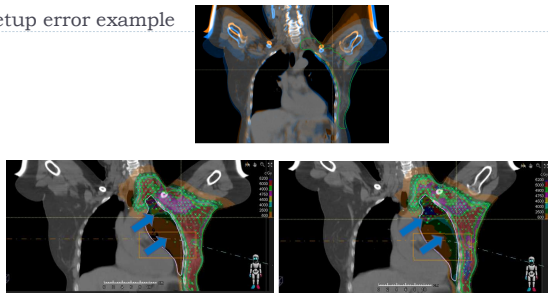
Ref: AAPM Task Group 176.

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Setup error example

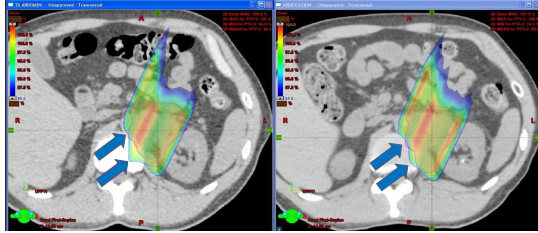


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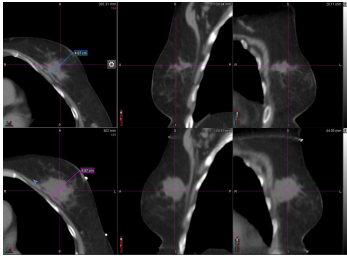
Range uncertainty due to physiological change such as small bowel filling change



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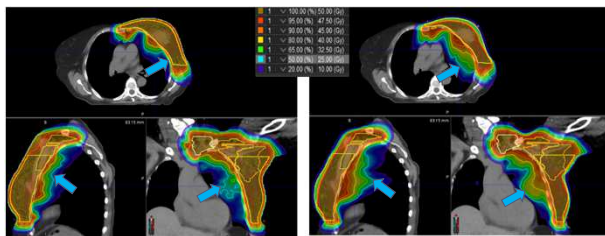
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Change of seroma example



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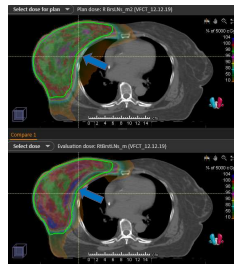
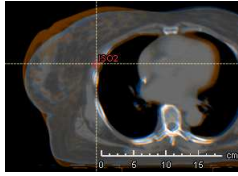
Dose distribution when with larger seroma

Dose distribution when seroma getting smaller.

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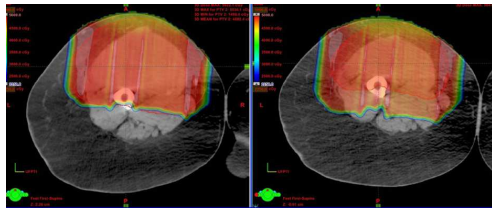
Edema example



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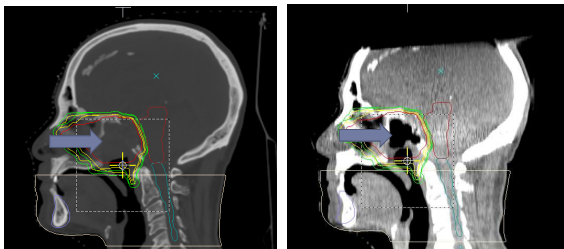
Tumor regression



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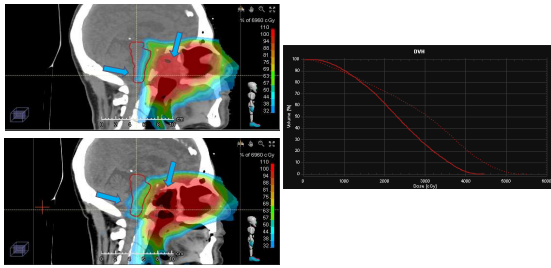
Tumor regression



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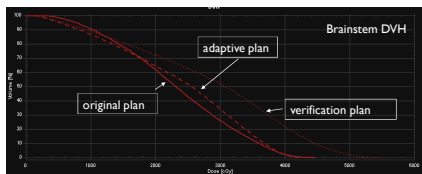
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Tumor regression



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Recommendations

- ▶ Include immobilization devices or patient support devices into the calculation if a proton beam goes through these devices.
- ▶ Robust treatment plan
 - ▶ Beam angle selection – minimize the potential WET variations.
 - ▶ If the proton beam angle cannot avoid potential WET variations, such as going through air cavities, larger range uncertainties should be allowed.
 - ▶ Appropriate treatment planning parameters and margins
 - ▶ Appropriate robust optimization settings
- ▶ Well defined IGRT protocols and residual setup error tolerance
- ▶ Closely monitoring the anatomy change and setup variations using CBCT
- ▶ Verification CT to assess the dosimetry stability
- ▶ Adaptive planning

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Immobilization device

- ▶ Uniform low density
- ▶ Homogeneous construction
For example, 'tennis racket' type table is not compatible with proton therapy
- ▶ Hardware clearance
- ▶ Range pullback accuracy

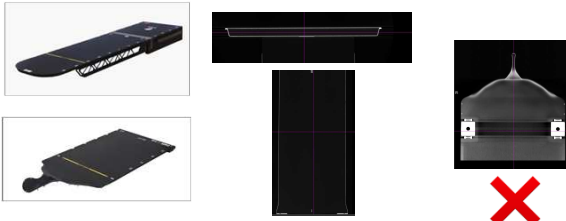
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Evaluate the homogeneity and internal structures

- ▶ Check inhomogeneities such as voids or regions of higher than nominal density.



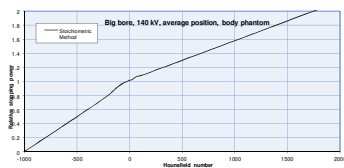
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HU to RSP calibration curve

- ▶ Using artificial tissue phantoms, the CT numbers are converted to relative proton stopping powers and then to WET for dose calculations.
- ▶ The materials used for immobilization device are usually different from the tissue materials used in the CT number calibration process, therefore, a measurement is required to check the accuracy of the stopping power calculation

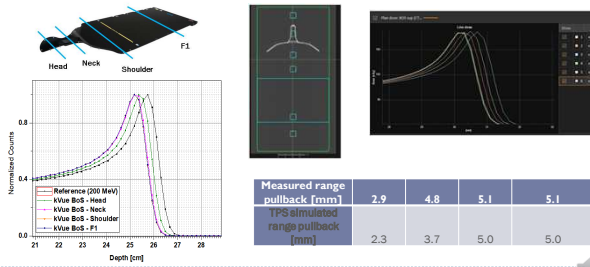


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Range pullback measurement on immobilization devices



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Characterize the immobilization device in TPS

- ▶ The change of proton distal range due to the immobilization device must be taken into account in the TPS. For proton radiotherapy, the couch top and immobilization devices in the beam path act as range shifters.
- ▶ If the measured range pullback matched the calculated one in TPS
 - ▶ Include the immobilization device in the external contour; incorporate it into the calculation.
- ▶ If the measured value does not matched the calculated one in TPS
 - ▶ Contour the immobilization device, override the CT number/physical density for the correct proton relative stopping power.

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HU override for non-biological tissue

- ▶ For example, breast implant
 - ▶ Saline-filled: 0.9% salt concentration distilled water
 - ▶ Silicone – polydimethylsiloxane (PDMS)

11%-17% RSP error

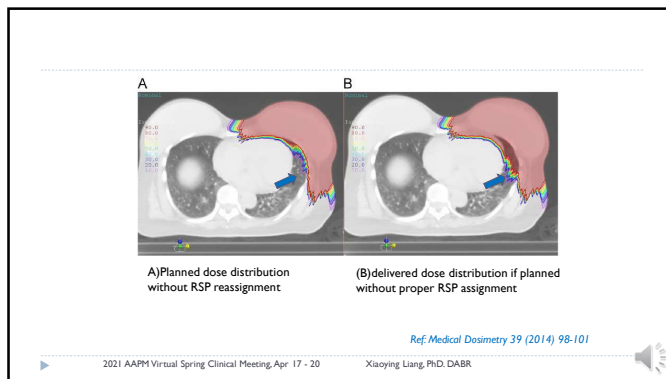
Table 2
Characteristics of test sample materials. Uncertainty values stated at 1 standard deviation level

Sample	Mass (g)	Volume from NCT (cm ³)	Measured density (g/cm ³)	Calculated Medical Dosimetry 39 (2014) 98-101	Measured	Converted pRLSP	Calculated pRLSP	Measured pRLSP
Distilled water	553.2 ± 0.4	505.00 ± 13.2	0.979 ± 0.023	1000	986 ± 4	0.994	1.000	n.p.
Tap water	950.6 ± 0.6	955.90 ± 19.0	0.994 ± 0.020	n.p.	987 ± 5	0.994	1.000	n.p.
Saline	403.4 ± 0.3	410.50 ± 10.7	0.983 ± 0.026	1010	1010 ± 3	1.010	0.998	n.p.
Sientra smooth	356.4 ± 0.3	369.68 ± 10.0	0.964 ± 0.030	1030	1123 ± 4	1.085	0.929	0.936 ± 0.016
Sientra textured	386.4 ± 0.3	399.65 ± 10.5	0.967 ± 0.025	1030	1121 ± 5	1.085	0.929	0.933 ± 0.015
Naturelle small	377.1 ± 0.3	391.52 ± 10.4	0.963 ± 0.026	1030	1120 ± 4	1.085	0.929	0.937 ± 0.016
Naturelle large	798.9 ± 0.6	826.20 ± 17.0	0.967 ± 0.020	1030	1110 ± 4	1.084	0.929	0.976 ± 0.014

Ref: Medical Dosimetry 39 (2014) 98-101

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Motion evaluation

- The purpose of assessment of the target motion –
 - Not only for margin,
 - Also for choosing the beam angle to minimize the effect of respiratory motion and interplay effect.
- For proton therapy, the dosimetric impact of tumor motion is a complex function of:
 - Inter- and intra-fractional motion
 - Treatment plan: beam angle selection, robust planning parameters
 - Delivery system: the scanning time and spot size,
 - Fractionation etc.

Ref: Chang et al. Int J Radiat Oncol Biol Phys. 2014; 90(4): 809–818.

A well defined procedure flow chart is recommended

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```

graph TD
    Start([Motion Analysis & Beam Angle Selection]) --> Calc1["STAD200 and STAD100  
Δ = [CWIPmean(%) - Wmean(%)]"]
    Calc1 --> Dec1{"Δ ≤ 3%"}
    Dec1 -- Y --> RPT1[RPT]
    Dec1 -- N --> Box1["50, 100% LRS, VRS"]
    Dec1 -- N --> Box2["R0/AD10"]
    Dec1 -- N --> Box3["LRS, VRS, BR, gating & others"]
    Box1 --> Calc2["V0/AD200 and STAD200  
Δ = [CWIPmean(%) - Wmean(%)]"]
    Box2 --> Calc2
    Box3 --> Calc2
    Calc2 --> Dec2{"Δ ≤ 3%"}
    Dec2 -- Y --> RPT1
    Dec2 -- N --> NoRPT([No RPT])
  
```

Example of Clinical workflow

Ref: Chang et al. Int J Radiat Oncol Biol Phys. 2017; 99(1): 41–49.

Range uncertainty

Source of range uncertainty in the patient	Range uncertainty without Monte Carlo	Range uncertainty with Monte Carlo
Independent of dose calculation:		
Measurement uncertainty in water for commissioning	± 0.7 mm	± 0.3 mm
Compensator design	± 0.2 mm	± 0.2 mm
Beam reproducibility	± 0.2 mm	± 0.2 mm
Pinpoint setup	± 0.7 mm	± 0.7 mm
Dose calculation:		
Biology (always positive)	± 0.3% #	± 0.3% #
CT imaging and calibration	± 0.8% ^a	± 0.5% ^b
CT conversion to tissue (excluding ^c -values)	± 0.3% ^c	± 0.2% ^d
CT grid size	± 0.3%	± 0.3%
Mean excitation energy (E-value) in tissue	± 1.5% ^e	± 1.5% ^e
Range degradation: complex inhomogeneities	± 0.7% ^f	± 0.1 %
Range degradation: local lateral inhomogeneities ^g	± 2.5% ^f	± 0.1 %
Total (excluding ^a - ^g)	2.7% ± 1.2 mm	2.4% ± 1.2 mm
Total (including ^a - ^g)	4.6% ± 1.2 mm	2.4% ± 1.2 mm

H. Paganetti, Phys. Med. Biol. 57, R99-R107 (2012).

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- Proton treatment planning needs to be done by experienced planners who understand the impact of range uncertainties
- Understand the sources that impact range uncertainties. Therefore, mitigate those that can be mitigated.
- Robust planning with appropriate range uncertainty margins

RO-ILS Case Study: Incorrect Density Factor

<https://www.astro.org/Patient-Care-and-Research/Patient-Safety/RO-ILS/RO-ILS-Education>

Case Example:

- › **Overview:** A typo error by a dosimetrist resulted in the PTV being assigned a density of 0. There was about 80% coverage on the PTV. This error was not appreciated by anyone as the dosimetrist finalized the plan.
- › **Details:** The evening before the scheduled start, the dosimetrist worked on transferring the approved plan to the oncology information system (OIS). The dosimetrist notified the medical physicist that the plan was ready for IMRT QA. Standard IMRT measurement-based QA was completed and did not identify the error. Physics was not aware the patient would be starting the next day and therefore did not perform the required second check of the treatment plan that evening. The following morning the plan still was not checked due to several scheduled special procedures and the lack of realization that the patient was starting that day. The treating therapists were very busy and failed to realize that the physics second check had not been performed, so the patient was treated according to the plan in place at the time. Physics then checked the plan in the evening after the patient's first delivered fraction. Upon this review, the physicist noticed that the PTV had been assigned a density of 0. Physics notified the dosimetrist and physician and the remaining 43 fractions were re-planned with the correct density, accounting for the dose already delivered during the first fraction. While there was only 80% coverage of the PTV for the first fraction, the subsequent re-planned fractions corrected for this deficiency and the patient continued treatment to completion.

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RO-ILS Case Study: Incorrect Density Factor

Contributing Factors/Root Causes:

- › Human error related to data entry (i.e., dosimetrist assigning the wrong density value to a treatment volume).
- › Rushed work and compressed timeline (e.g., dosimetrist working on plan the evening before the patient started treatment).
- › Numerous special procedures scheduled on the same morning, preventing the physicist from performing the second check as part of normal work duties (e.g., as they would have done even in a non-rushed situation).
- › Ineffective communication of scheduled treatment start date.
- › Lack of awareness of the ineffective communication of scheduled treatment start date.
- › Failure of staff to perform established process (e.g., failure of busy therapist staff to perform comprehensive initial chart check, assessing all pertinent information that needs to be completed prior to patient start treatment such as, approval of the prescription, signed consent form and verifying physics QA was completed).
- › Lack of a forcing function (e.g., hard stop) to prevent treatment of patient in the absence of a completed second check.

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Thank you!