



Understanding IMPT and Delivery Uncertainties

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Session Objectives

- Review of treatment safety margins and how they relate to proton therapy
- Discuss IMPT Optimization methods
- Introduce the concept of "Robustness"
- Define intra-treatment concerns specifically important to IMPT

An important statement from the AAPM:

The essential responsibility of the Medical Physicist to assure the safe, effective and consistent delivery of radiation*

Acceptance Testing

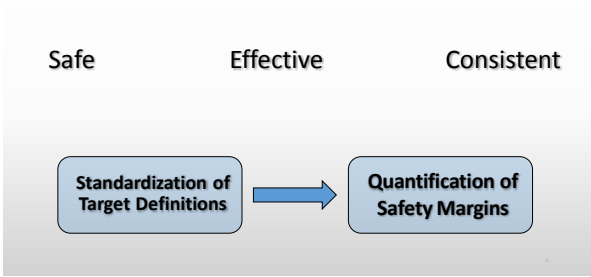
Quality Assurance

Commissioning

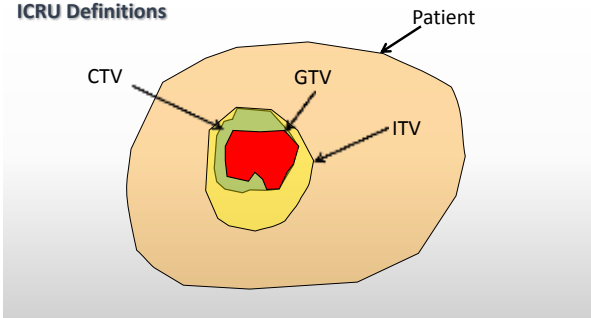
Procedural Guidelines

* AAPM Scope of Practice for a Qualified Medical Physicist

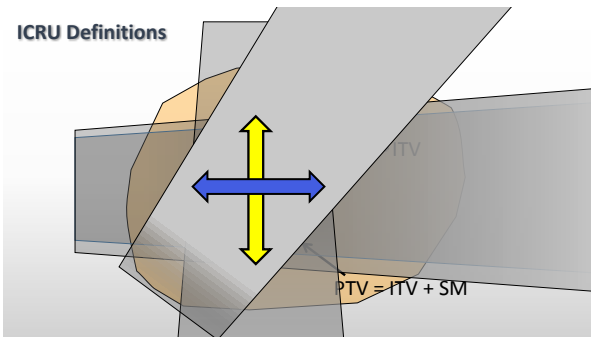
With Regards to Treatment Planning.....



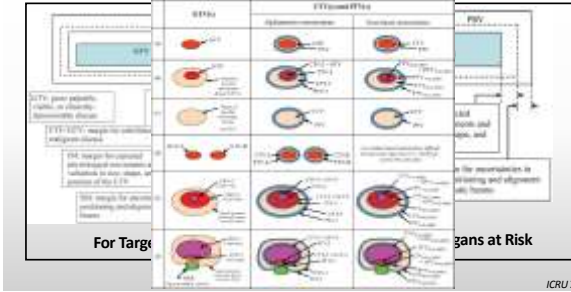
ICRU Definitions



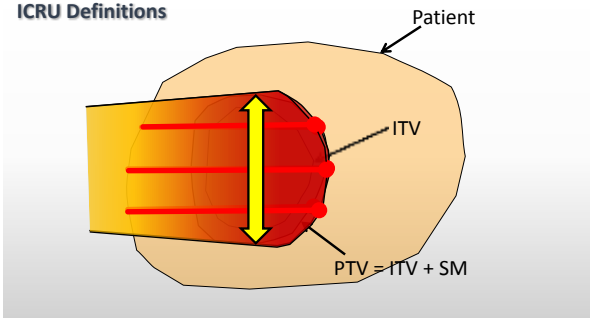
ICRU Definitions



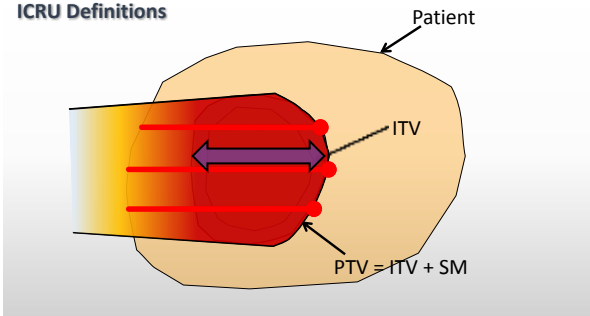
ICRU : Safety Methods to Avoid a Geometric Miss of the Target



ICRU Definitions



ICRU Definitions



Safe Effective Consistent

Are there other safety considerations that need to be considered for the distal edge margins??

Proton Range Uncertainties

Horizontal lines for taking notes.

With Protons : We must always consider for Range Uncertainty

Table 1. Summary of uncertainties in the range of protons in water and soft tissue.

Source of uncertainty	Uncertainty (cm)	Relative uncertainty (%)
Stochastic fluctuations in ionization cross-sections	0.01-0.02	0.1-0.2
Stochastic fluctuations in ionization cross-sections (Monte Carlo)	0.01-0.02	0.1-0.2
Stochastic fluctuations in ionization cross-sections (Analytical)	0.01-0.02	0.1-0.2
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Table 2. Summary of uncertainties in the range of protons in water and soft tissue.

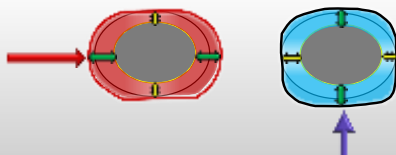
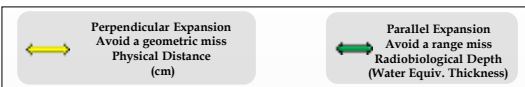
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Stochastic fluctuations in ionization cross-sections (Analytical)	0.01-0.02	0.1-0.2

Table 3. Summary of uncertainties in the range of protons in water and soft tissue.

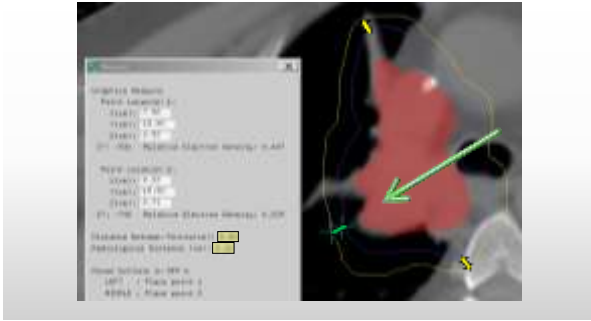
Source of uncertainty	Uncertainty (cm)	Relative uncertainty (%)
Stochastic fluctuations in ionization cross-sections	0.01-0.02	0.1-0.2
Stochastic fluctuations in ionization cross-sections (Monte Carlo)	0.01-0.02	0.1-0.2
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Horizontal lines for taking notes.

Margins for Setup and Range Uncertainty with Protons



Horizontal lines for taking notes.

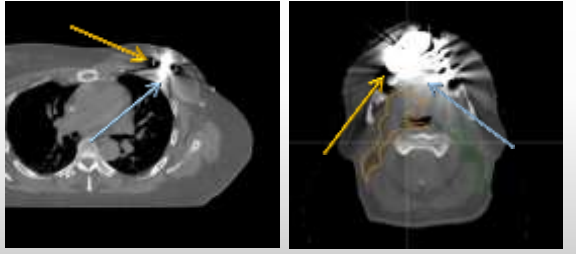




Using a CT as the Patient “Map” is an Area of Specific Concern for Protons

- Trying to make our CT scanner a spectrometer
 - Two tissues can have same HU but different RSP
- CT scans are not perfect
 - Noise
 - Beam hardening
- Stoichiometric Method is valid for tissue type materials. Anything not natural can have large errors.
 - Contrast
 - Fillings
 - Implants

Artifact caused by the Filling Port of a Chestwall Expander / Dental Fillings



HU to RSP Conversion Errors in a Silicone Breast Prosthesis

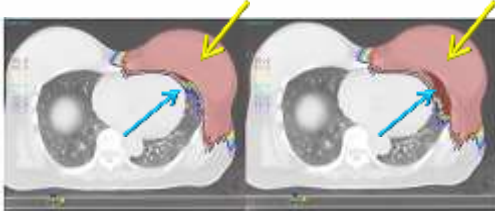


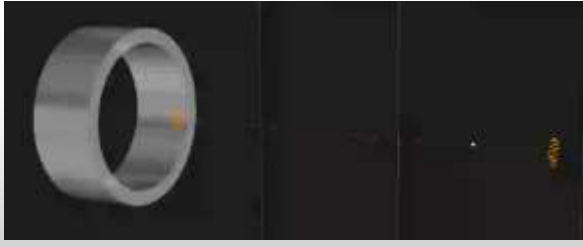
Fig. 3: Treatment plan for patient with silicone breast prosthesis. (a) Planned dose distribution without RLSP reassignment. (b) Delivered dose distribution if planned without proper RLSP assignment.

Mayers et al. Med Dosim. 2014 Spring;39(1):98-101.

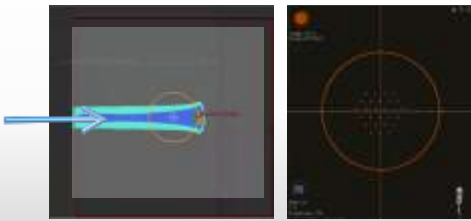
Delivery of a Sphere of Dose using PBS



Delivery of a Sphere of Dose using PBS



Spot / Layer Patterns for a Sphere



Dose Distribution

Beam's Eye View

How can we obtain these complex, 3-D spot patterns?

Inverse planning techniques

- Iterative minimization of an objective function.
 - Cover target areas
 - Minimize dose to organs at risk
- Exceptional potential for computing and mathematical "tricks"
 - Minimizing the effects of positional errors and range errors directly into the cost function (Robust Optimization)
 - Including effects of motion into the cost function (4-D Optimization)
 - Multi Criteria Optimization (MCO)
- Two different optimization methods used to guide the objective function



Commonly used IMPT Optimization Methods

Single Field Optimization (SFO)

Uniform Dose is delivered to the entire target by each field individually

Less sparing of critical structures

Less sensitive to Set-up/Range errors

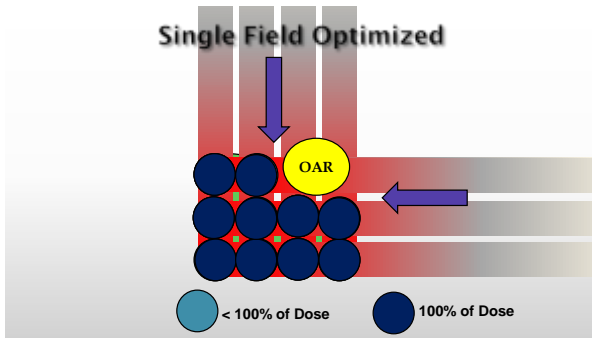
Multi Field Optimization (MFO)

Spot weights of all fields are optimized together. The spot weight of one field will rely on another field's dose to create an integrated uniform target dose

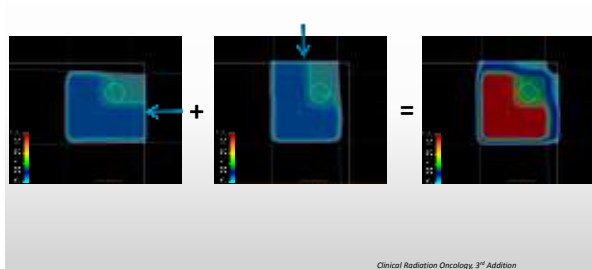
Better for sparing critical structures

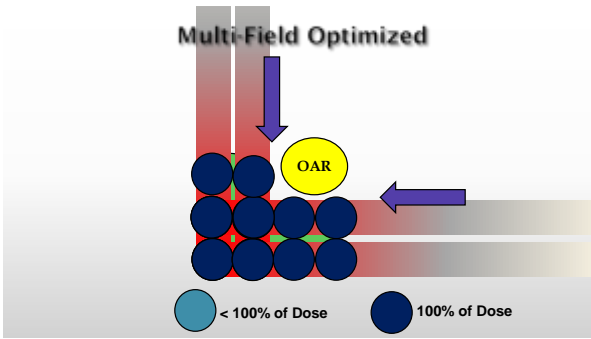
More sensitive to Set-up/Range errors

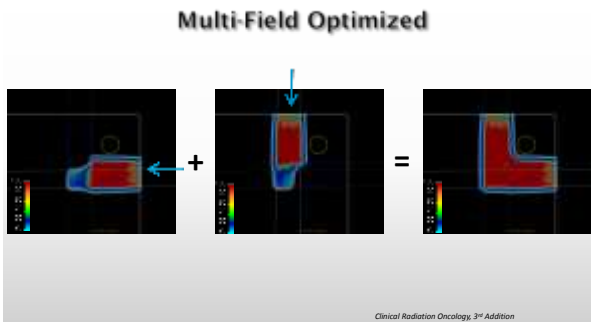
Single Field Optimized

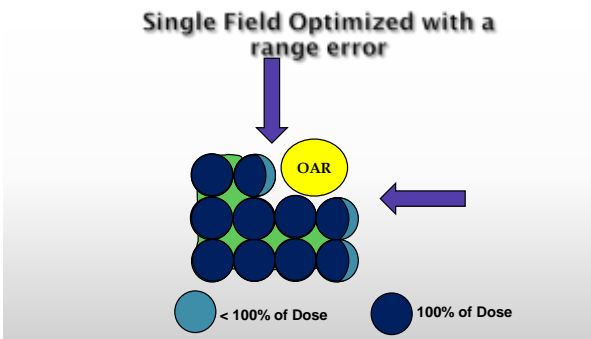


Single Field Optimized

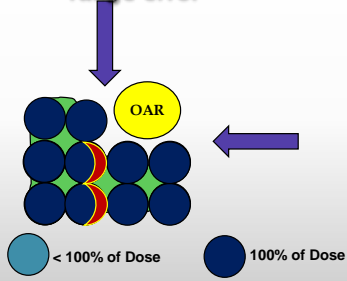








Multi Field Optimized with a range error



The physicist is tasked to

Quantify the effects of:

- Non-ideal set-up
- Intra-fraction motion
 - Respiratory motion

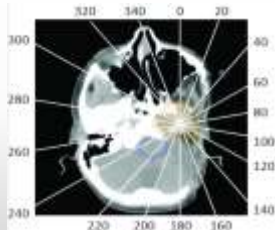
“Robustness”

- Range uncertainty
- Inter-fraction motion
 - Anatomical consistency

Prospective Robust Planning

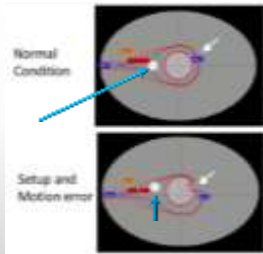
Beam Angle Optimization prior to Spot optimization

- Evaluate path-lengths and the effect of range error and set-up uncertainties
- Concept can be expanded to 4-D evaluations



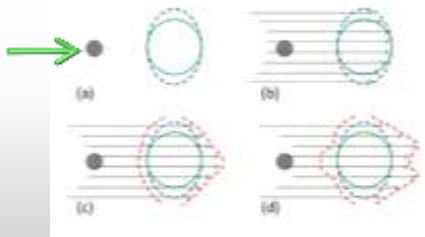
Cao et.al Med Phys. 2012 Aug; 39(8): 5248–5256. MDA

The Effect of Set-up Errors on SFO Plan



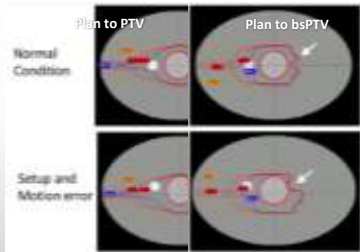
Park et al. Int J Radiat Oncol Biol Phys. 2012

For SFO: Beam Specific PTV : bs(PTV)



P. Park et al Int J Radiat Oncol Biol Phys. 2012

The Effect of Set-up Errors on SFO Plan



Park et al. Int J Radiat Oncol Biol Phys. 2012

What about MFO methods?

Robust Optimization

- Add penalties into the cost function for robustness
- Allow the planning system to score robustness on a spot to spot basis AND how one spot will effect the overall sensitivity to potential plan degradation.
- Spots with "poor" robustness (high sensitivity to plan degradation) will be penalized by iteratively decreasing, and potentially, eliminating their intensity
- There is potential for LET / Biological Dose Optimization

Robust Optimization

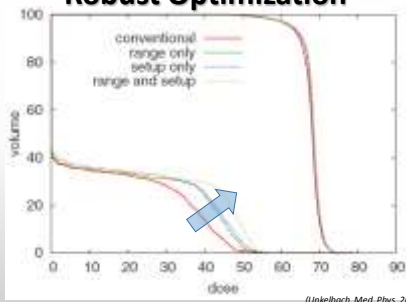
• Range Uncertainties and Set-up Errors

- 1) Probabilistic approach
 - Range and positional errors are parameterized and incorporated into the objective function
 - spot range/position/weight is a random variable
- 2) Optimize on "worst case" scenarios



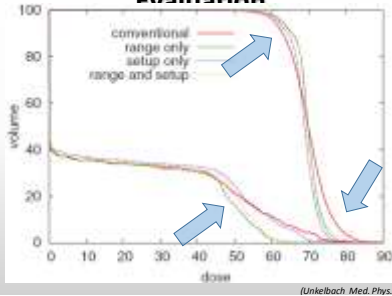
Unkelbach Phys. Med. Bio. 2007, Med Phys 2009

Robust Optimization



(Unkelbach, Med. Phys. 2009)

Robust Optimization Evaluation

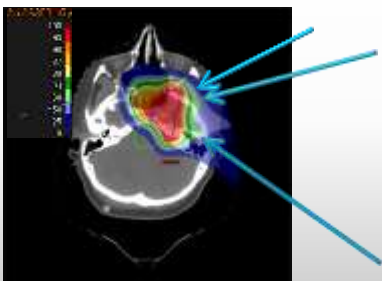


(Unkelbach Med. Phys. 2009)

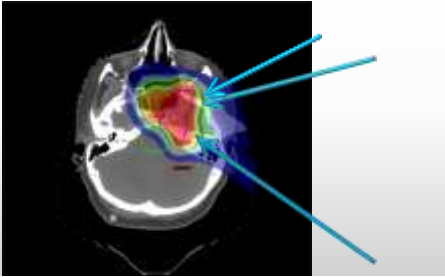
Planned with optimization



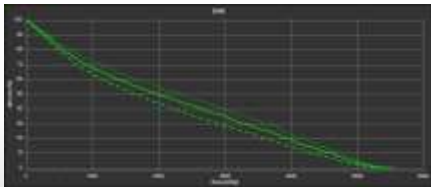
Robust Optimization for OAR



Robust Optimization for OAR



Robust Optimization for OAR



Dose	RO/PO	Clinical goal	Value
Plan dose: To 41 G...	Roar/tem	All mean 5700 uGy dose at 0.0 cm ³ volume	5700 uGy
Perturbed dose: CT...	Roar/tem	All mean 5700 uGy dose at 0.0 cm ³ volume	5700 uGy
Perturbed dose: CT...	Roar/tem	All mean 5700 uGy dose at 0.0 cm ³ volume	5700 uGy

The balancing act of trade-offs



- Multi-Field Optimized ————— Single Field Optimized
- Target Coverage ————— Critical Organ Dose
- High Quality plan ————— Robustness

Degeneracy within IMPT plans Which is best?

- Multi-Criteria Optimization MCO

- Database of plans each emphasizing different planning objectives are pre-calculated to approximate the Pareto surface
- Pareto surfaces are navigated to find the optimal balance
- Range and Set-up uncertainties can be included



Chen et al. Phys. Med. Biol. 2012

The physicist is tasked to

“Robustness”

Quantify the effects of:

- Non-ideal set-up
- Intra-fraction motion
 - Respiratory motion

Robustness Evaluation

- Range uncertainty
- Inter-fraction motion
 - Anatomical consistency

Important note : Robust Optimization does not guarantee a robust plan

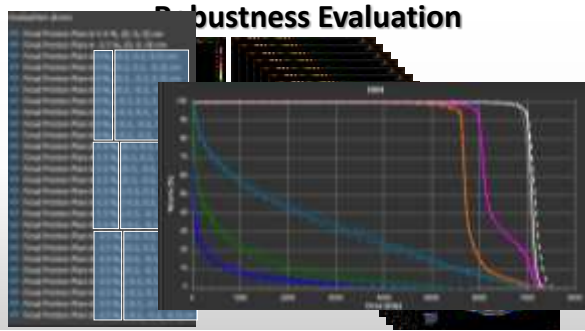
Robustness Evaluation

Quantify the differences in quality between the planned and the delivered dose in the presence of uncertainties

Robust Plan Evaluation includes :

- Calculation and Evaluation of many “Worst case” scenarios
 - Systematic offset of HU conversion (-3.5% , + 3.5%)
 - Systematic offset of set-up error (x= +/- 5mm, y= +/- 5mm, z= +/- 5mm)

Lamax Phys. Med. Biol 2008



The concept of “Robust Evaluation” really has limitations

- Impossible to look at all potential scenarios
- In reality there is a combinations of Random AND Systematic errors
 - Set-up errors are random
 - Range errors are systematic
 - Beam Hardening
 - Tissue Dependent
 - Range errors are NOT uniformly distributed
- It is essential to have clear communication between physics and physicians of potential target coverage limitations and possible OAR doses.
- Only as good as the patient model that you give it.
 - What if the patient changes.....

Adaptive Planning



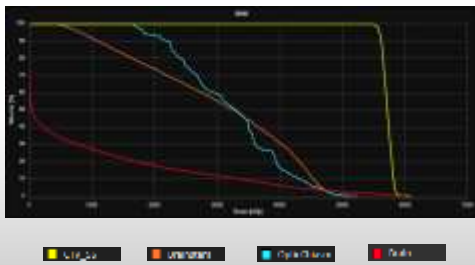
All radiation treatment delivery is sensitive to anatomical changes

- Especially in particle therapy, a clear understanding of the magnitude and potential consequences of any anatomical change needs to be defined
- IMPT delivery, especially MFO optimized plans , can be very sensitive to anatomical changes.
- Workflows need to be in place to detect, evaluate and correct unexpected anatomical changes.

Adaptive Planning : Naso-pharynx to 56Gy_(RBE)



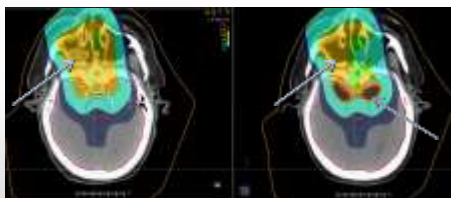
Approved Plan : Naso-pharynx



Large Change in Target Region Anatomy



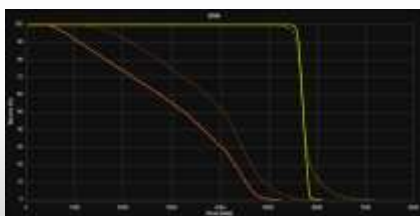
Calculation of Initial Plan on New Image Set



Initial Plan

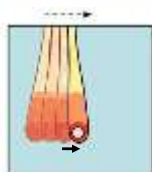
On Treatment Evaluation

DVH with modified anatomy



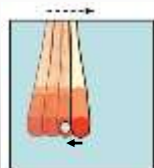
Legend: Yellow, Blue, Red, Orange

Interplay Effects Due to Motion and Delivery Timing



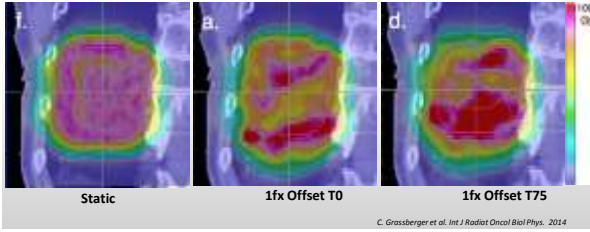
Motion in same direction = Dose Increase

Motion in Opposite direction = Dose Decrease



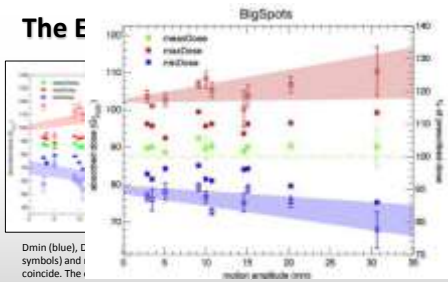
ICRU 78

IMPT delivery has a time structure The Effects of Respiratory Motion



C. Grassberger et al. Int J Radiat Oncol Biol Phys. 2014

The E



Dmin (blue), E symbols) and coincide. The shaded areas represent a linear fit through the maximum and minimum values of to guide the eye.

C. Grassberger et al. Int J Radiat Oncol Biol Phys. 2014

Daily Under-dosing and 2y-LC

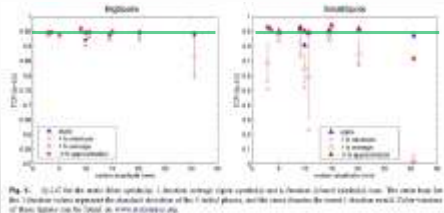


Fig. 4. 2y-LC for the same data conditions. 1. Scatter through figure symbols and 2. Linear fit through symbols. The color bar for the 2y-LC values represent the absolute distance of the 2y-LC values, and the color bar for the 2y-LC values represent the absolute distance of the 2y-LC values. Color values of 0.00, 0.05, 0.10, 0.15, 0.20, 0.25, 0.30, 0.35, 0.40, 0.45, 0.50, 0.55, 0.60, 0.65, 0.70, 0.75, 0.80, 0.85, 0.90, 0.95, 1.00.

C. Grassberger et al. Int J Radiat Oncol Biol Phys. 2014

Motion effects mitigation

Treatment Delivery

- Repainting
- Gating
- Breath Hold / Compression
- Beam Tracking

Treatment Planning

- Planning Methods
 - Using more Fields
 - More fractions (>5)
- 4-D Robust Optimizations
- 4-D Dose Evaluation

Strategies for addressing Motion : Repainting

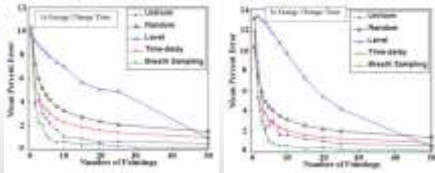


Figure 4. Number of paintings required to reduce the dosimetric effect of motion (breathing period: 4.4 s, motion trajectory: sin4, 30 fractions), 1 s and 5 s energy change time.

J. Seca et al. Phys. Med. Biol. (2009)

In Summary

- IMPT is an essential tool in the effort to reach the full potential of particle planning. IMPT methods generate better plans and to more treatment sites.
- IMPT methods are sensitive to range errors, set-up errors, motion and anatomical variances. (Robustness)
- Robust Optimization methods can be used to minimize these sensitive effects along with Robust Evaluations to quantify the consequences.
- IMPT planning is a trade off of many variables. Careful planning with effective communication is necessary between the physics and physician teams.

The image shows the words "THANK YOU" in a bold, 3D, red font. The letters are thick and have a slight shadow underneath, giving them a three-dimensional appearance. The text is set against a white background with a subtle gradient.
