

To promote excellency in patient care and innovative proton treatment

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Uncertainties in the clinical practice

# Nominal



# Delivered on a single fraction





There are different kind of uncertainties:

- a) Density uncertainties
- b) Range uncertainties
- c) Set up uncertainties

# d) RBE

e) Beam delivery



They can appear in different step of the entire radiotherapy workflow: from planning CT to the dose delivery.





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## Inherent CT uncertainties: calibration from HU to RPSP



~1% range uncertainties through soft tissues, ~2% through bone ~1.5mm range uncertainty in brain, ~3.5mm for prostate (lateral beams)

<sup>1</sup>Schneider et al. 1996, <sup>2</sup>Schaffner and Pedroni 1998

### CT artifacts can compromise CT data quality

Artefacts along the spinal cord due to titanium implants



Artefacts due to metal implants in the Teeth (normally not titanium)



### CT artifacts : how we can deal with them

CT data set with artefacts

Manual delineation of the soft tissue areas, HU substituted



Additionally: beam hardening effect, noise and resolution. Planning CT errors: they are systematic and contribute to range uncertainties.

#### Single field trough the metal implant: extreme case





## Stopping power profiles

#### Single field trough the metal implant: extreme case





They can appear in different step of the entire radiotherapy workflow: from planning CT to the dose delivery.





Planning

#### Density heterogeneities: degradation of the Bragg peak shape



Protons Through Base of Skull: 90 to 20% fall of increases from 6 to 32 mm

Urie et al, PMB, 1986, 31;1-15



# Example field through relatively homogenous anatomy



# Example field through very inhomogenous anatomy



Lomax PMB 2008a



When planning it is important to select field directions which go trough the minimum possible in-homogeneities.

They play also a role when we come to the positioning process







They can appear in different step of the entire radiotherapy workflow: from planning CT to the dose delivery.



Planning



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SFUD



SFUD plans are generally more robust than the IMPT plans (highly inhomogeneous individual dose distribution are 'patched' inside the target volume)



57

50

40

30

20

10

5

Evaluating uncertainties: max to min dose distributions

This indicates the maximum differences (~ error bars) expected in the plan due to set up errors (Albertini F. et al. Phys Med Biol. 2011) **SFUD** IMPT



- SFUD more robust than IMPT in the target
- IMPT plan more robust for brainstem

Lowe et al "Incorporating the effect of fractionation in evaluating the robustness of proton plans to set-up uncertainties"

Margins increase robustness for SFUD, but not necessarly for IMPT.



They can appear in different step of the entire radiotherapy workflow: from planning CT to the dose delivery.





Daily positioning protocol significantly reduce patient misplacement







"water equivalent" patient

### Set up errors combined with density heterogeneities

#### patient geometry



Even after image guided re-positioning, density heterogeneities can somewhat degrade the delivered dose due to residual positioning errors



They can appear in different step of the entire radiotherapy workflow: from planning CT to the dose delivery.





#### Nasal cavity fillings



Every week the repeated CT slices acquired showed a significant different changing in the cavity fillings, with consequences on the delivered dose distributions.



#### Nasal cavity fillings

Nominal dose distribution (CTV in green, spinal cord in red)



Recalculated dose distribution



Placidi et al. "The effect of anatomical changes on PBS proton dose distributions: a retrospective review of 951 patients treated at PSI"



## **During treatment**

#### Weight changes

3 fields IMPT plan, pt lost 1.5 kg.



Cauda equina, included in the PTV was spared







#### Volume shrinkage (neck lymph nodes)



Node shrinkage of about 1 cm; It was detected during the course of treatment.

Replanning was done due to the overdosage







They can appear in different step of the entire radiotherapy workflow: from planning CT to the dose delivery.





# **Delivered dose**



U position dynamic correction (teaching) [cm]



Meier G. et al. Independent dose calculations for commissioning, quality assurance and dose reconstruction of PBS proton therapy. Phys Med. Biol 2015 Apr 7; 60(7): 2819-36



Scandurra et el."Is what we plan what we actually deliver? Using machine log-files to analyse delivery of 20 clinical PBS plans on PSI Gantry 2". Poster at PTCOG 54



Different ways to deal with the (range and set up) uncertainties:

1. patient monitoring (detect range differences as soon as possible ideally daily)

2. adaptive therapy (adapt the plan, as soon as possible ideally daily)

3. robust planning

(reduce a-priori the impact of range uncertainties)

# Dealing with uncertainties





Dealing with uncertainties

## 1. Range probe



# Single pencil beam with going trough the patient residual range measured in the MLIC

PhD work of Abdel Hammi (PSI)



Dealing with uncertainties

## 1. Range probe 2D

#### 2D RP 5x5mm spacing.



# Residual positioning error and anatomical changes



PhD work of Abdel Hammi (PSI)



# Dealing with uncertainties





# 3. Robust planning

- 1. Careful selection of field incidences and planning techniques (as IMPT and SFUD)
- 1. Automatic incorporation of all the errors (range, set-up) in the optimization process (change of the cost-function)
- Unkelbach J et al 2009 Med Phys. ; Unkelbach J et al 2007 PMB; Maleike, Flynn (Ex Raysearch)
- Changing the optimization starting condition:

   a. manual selection of beam angles avoiding or
   penalizing path going through sensitive areas
   b. changing the initial beamlet fluences

Lomax A et al, 2001 Med Phys; Albertini F et al, 2010 PMB

There are different kind of uncertainties (i.e. heterogeneities, positioning, range, delivery, RBE...)

They come into play during the entire workflow

Different possible solutions are based on:

- a. Image guidance
- b. Daily adaptive
- c. Robust optimisation

The simplest and cheapest solution is based on: *Multiple fields with optimised directions* 



# Thank you!

# ...a special thank to Tony Lomax , Francesca Albertini and all my CPT colleagues



Uncertaities in the clinical practice:

Head and neck cases very challenging due to many concomitant problems:

-denti, artefatti, eterogeneneità patient positioning, shrinkage of the nodes...(satangunantan)

- -Organ motion (slide di Tony..ma questo merita talk a parte)
- -Referenze a poster di Dan e gabriel (eventuale slide su log file calculation)

Differenza di uncertainties in SFUD/IMPT!!!!!! Sostituire range adaptive con plan adaptation

Immagine su artefatti denti, eventualemnte tagliare ancora una slide di Stefania. Summary

#### Daily process Daily 3D concept... (3D) imaging Pre-treatment Error in spot position Define field Field geometry geometry Accumulated dose CT/MR Manual correction of magnet settingsl Reference CT/MR Accumulate dose 3D imaging Spot number Log files Reconstruct delivered dose Reconstruct ed dose

# Dynamically Adapted Radiotherapy (DART)