### INTER-FRACTIONAL MONITORING

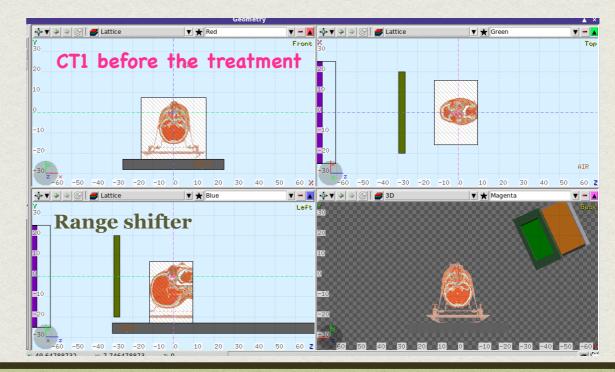
ARPG metting 02/05/19
Marta Fischetti

The DP capability to spot the inter-fractional changes (during the treatment) in the dose deposition, using the charged fragments emission shape (POCA), has been investigated with a Monte Carlo simulation using

the FLUKA software:

 Two CT: before the treatment and after the toxicity onset

- Same TP for each CT
- Real Positioning
- One fraction of <sup>12</sup>C ions

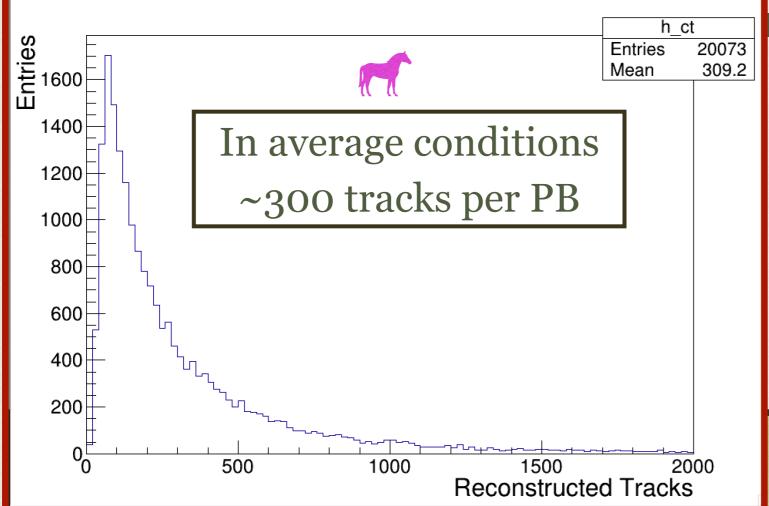


- We don't need to unfold the "matter effect"
- We have used the '1D' projections along the PB direction to perform a quantitative comparison -> Kolmogorv and  $\chi^2$  tests
- Low statistics for single PB ( $^{\sim}300$  tracks in most populated bins): Packing PB->5x5x3 = 75 PB (Volume = 1cm x 1cm x 6mm)

The DP capability to spot the inter-fractional changes (during the

treatment) in the dose deposition using the charged fragments emission







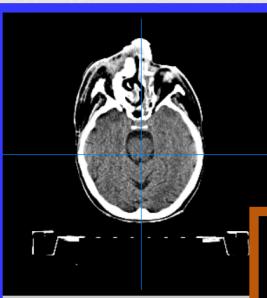
Plots marked with a pink horse are in preparation for PMB paper

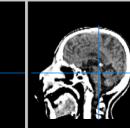
perform a quantitative comparison

- ->Kolmogorv and X²
- Low statistics for single PB ( $^{\sim}300$  tracks in most populated bins): Packing PB->5x5x3 = 75 PB

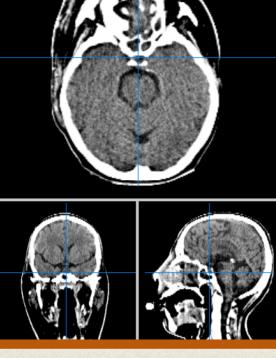
 $(Volume = 1cm \times 1cm \times 6mm)$ 

#### CT1 before the treatment

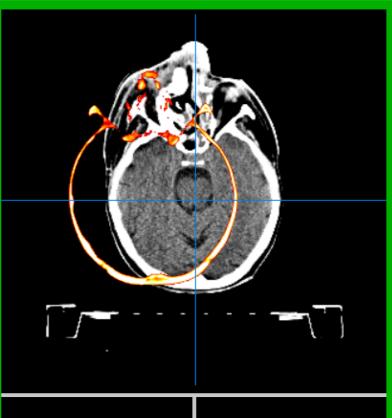


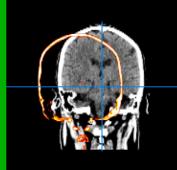


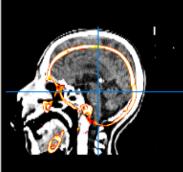
CT2 after the toxicity onset



CT2 overlaid to CT1



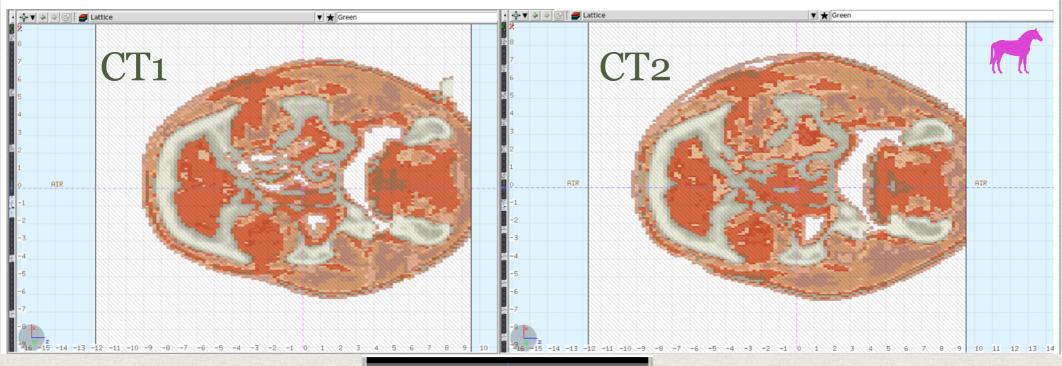




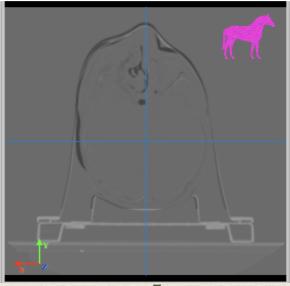
We have to align the two CT to use the same TP

Manual alignment isn't enough accurate

We solved the alignment problem of the two CT using flirt software!!!!



The subtraction
between CT1 and CT2 is
shown in gray scale



GOOD agreement

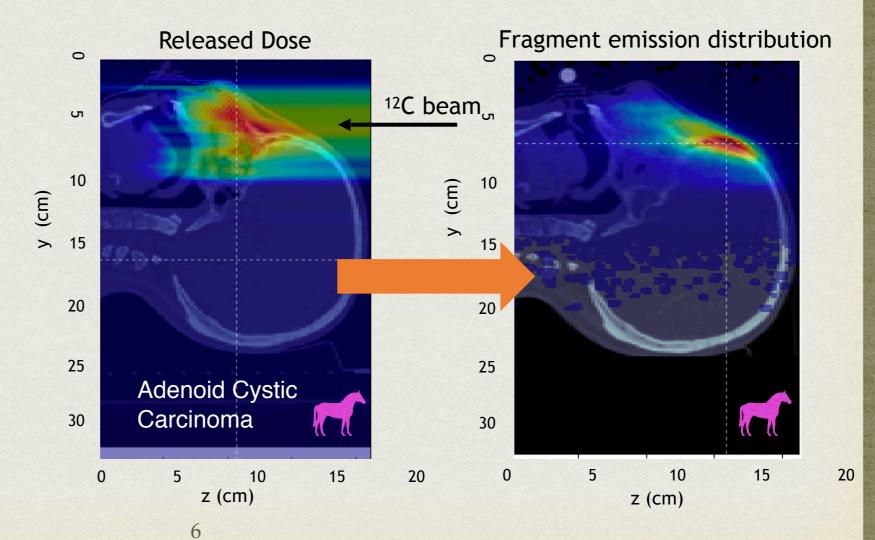
This treatment is composed by 3 fields: B1, B2, B3.

Firstly I analyzed B3:

the best condition for us

Exit window Range shifter 80 -70 -60 -50 -40 -30 -20 -10 0

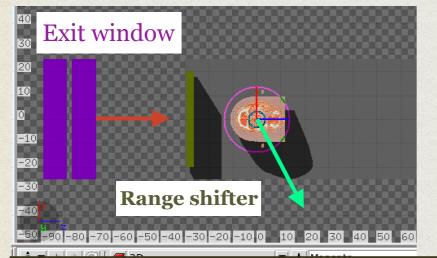
Fragments are mostly produced at the entrance point inside the patient and are absorbed by the patient body in their exit path towards the detector

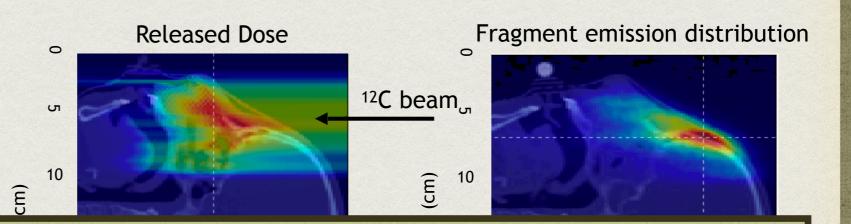


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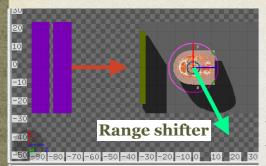
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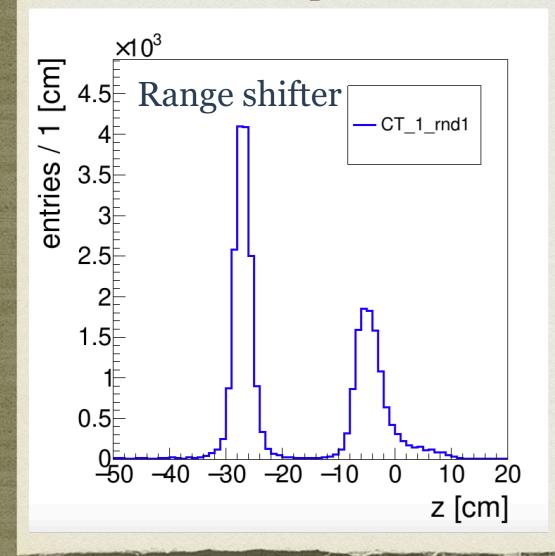


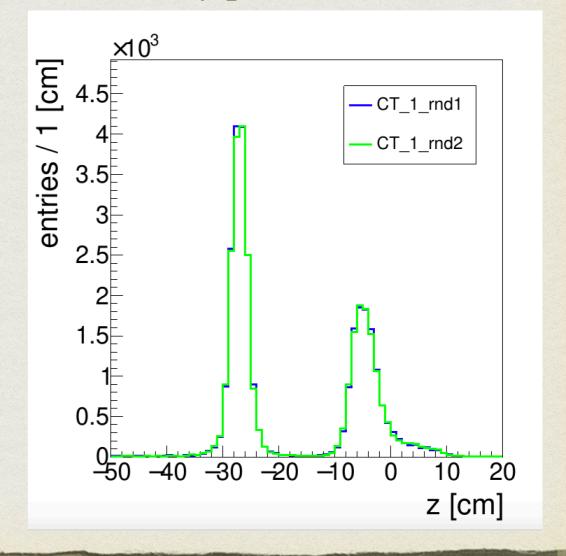
- Inter-fractional monitoring in dose deposition is done using the '1D' projections of secondary fragments emission vertex (POCA) along the PB direction
- More detailed method of '3D' comparison will be studied soon

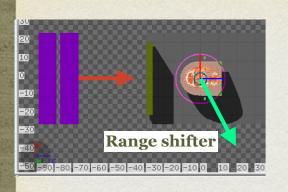
20



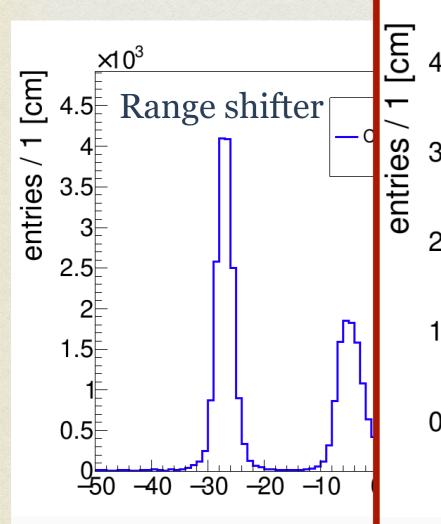
Reproducibility study of the method was done producing the same MC simulation (using the same CT1 scan and the same treatment plan) with different random seeds and comparing, super PB per super PB, the resulting profiles of secondary particles

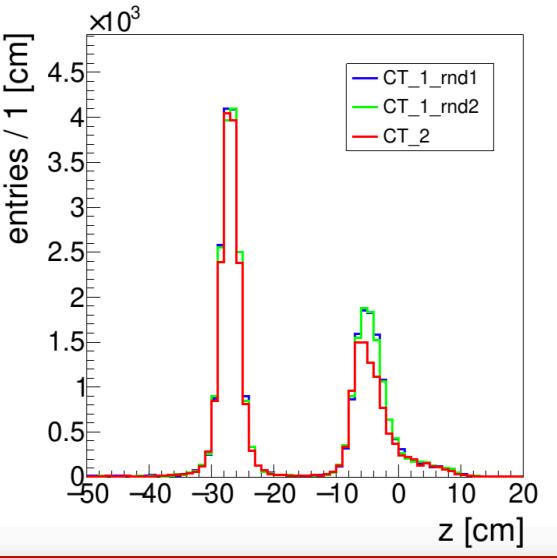


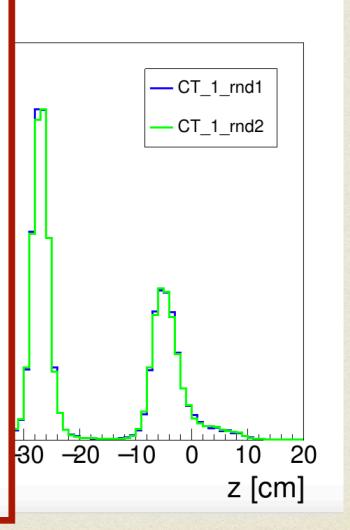


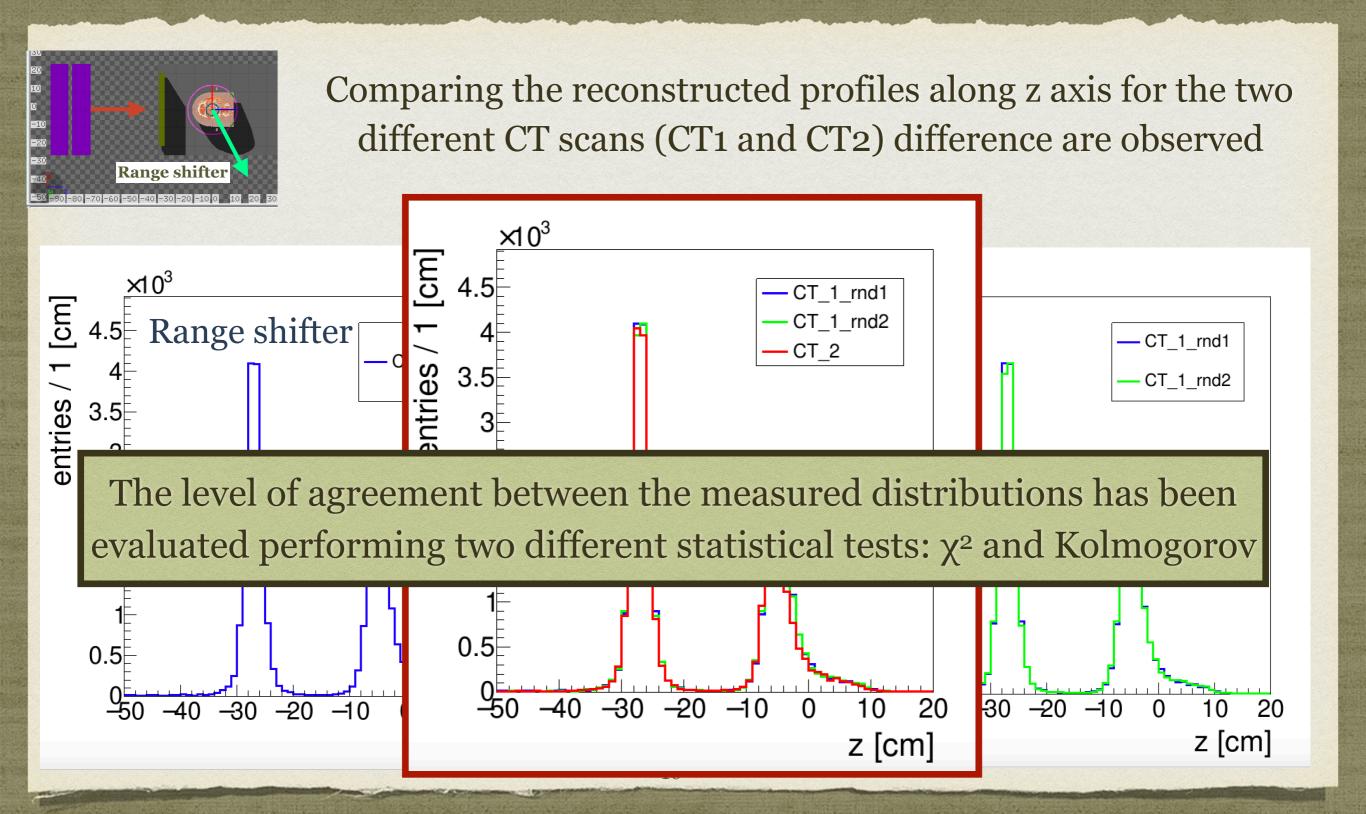


Comparing the reconstructed profiles along z axis for the two different CT scans (CT1 and CT2) difference are observed







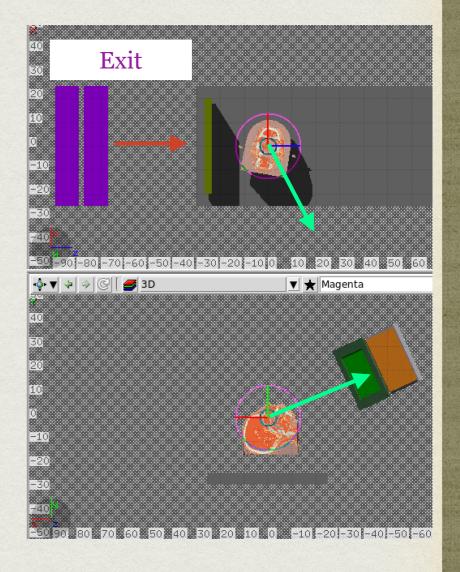


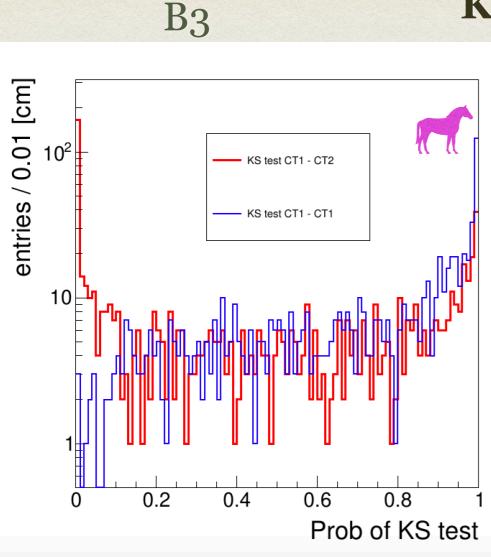
Firstly I analyzed B3: the best condition for us

Exit window ▼ ★ Magenta Secondly I analyze also B2

Another test in a worst condition has been done

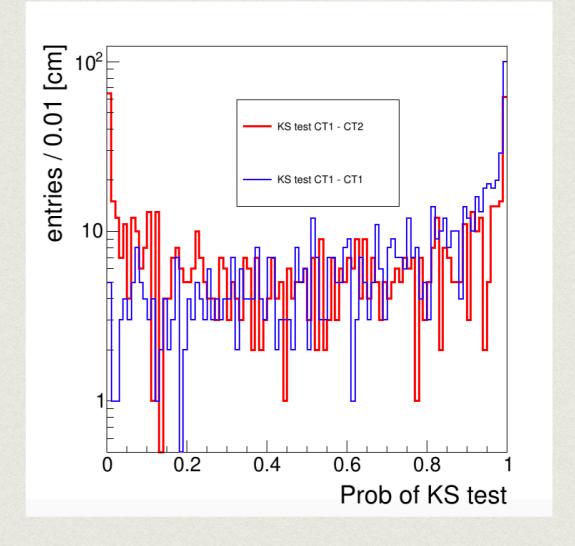
Fragments have to travel a bigger path inside the body before reaching the detector







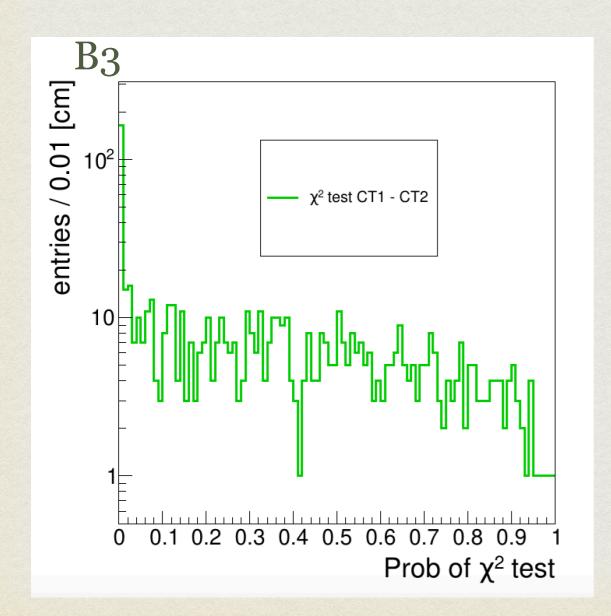




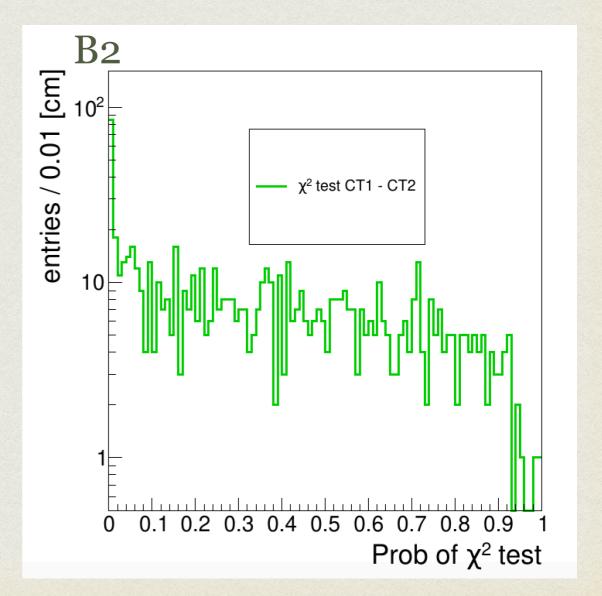
When CT1 and CT2 are compared (red), there's a clear evidence that in some superPB there's no agreement btw the measured distributions. Instead, when just checking the statistical fluctuations (blue) such peak at low p(KS) disappears

 $\chi^2$ 

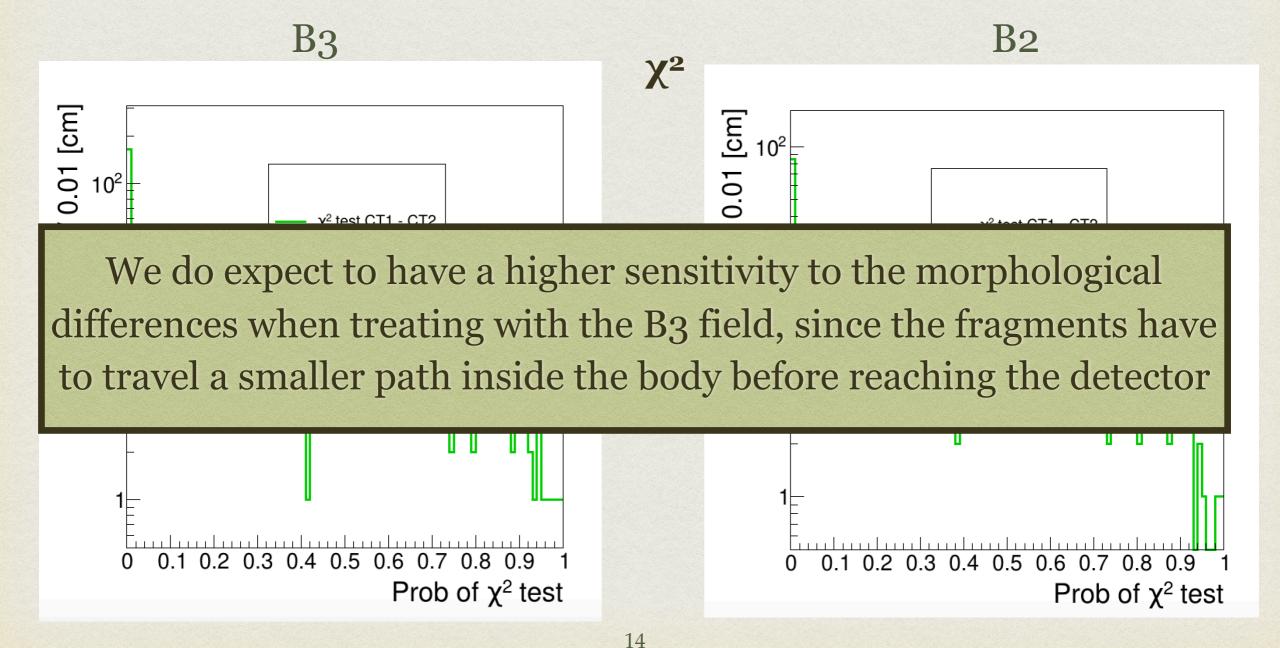
The χ² test was also studied because the results provided are binning dependent



Kolmogorov test are bin independent. It compares the cumulative of the distribution

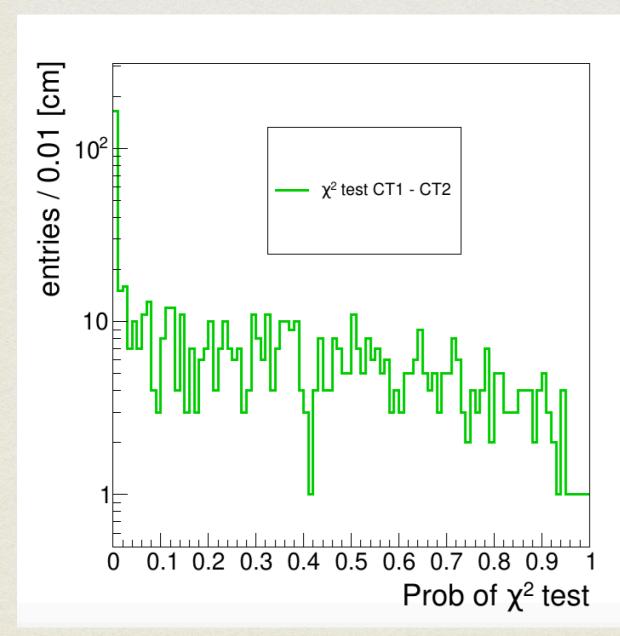


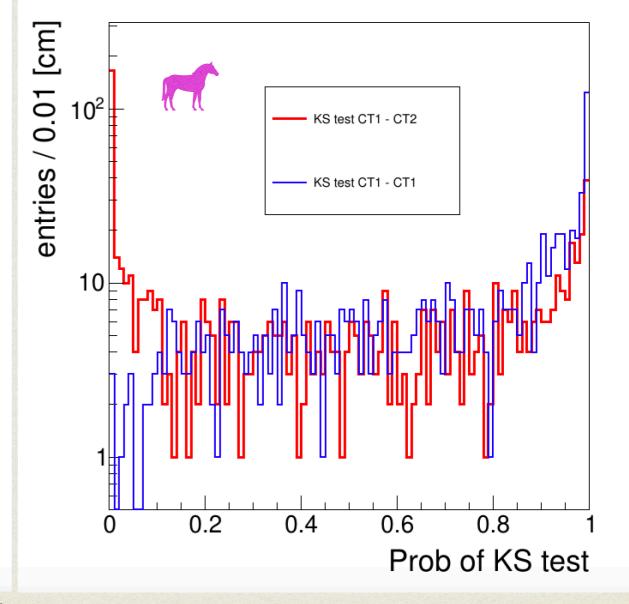
The  $\chi^2$  test was also studied as a further test to verify the robustness of the results provided by the statistical analysis done with the kolmogorov test



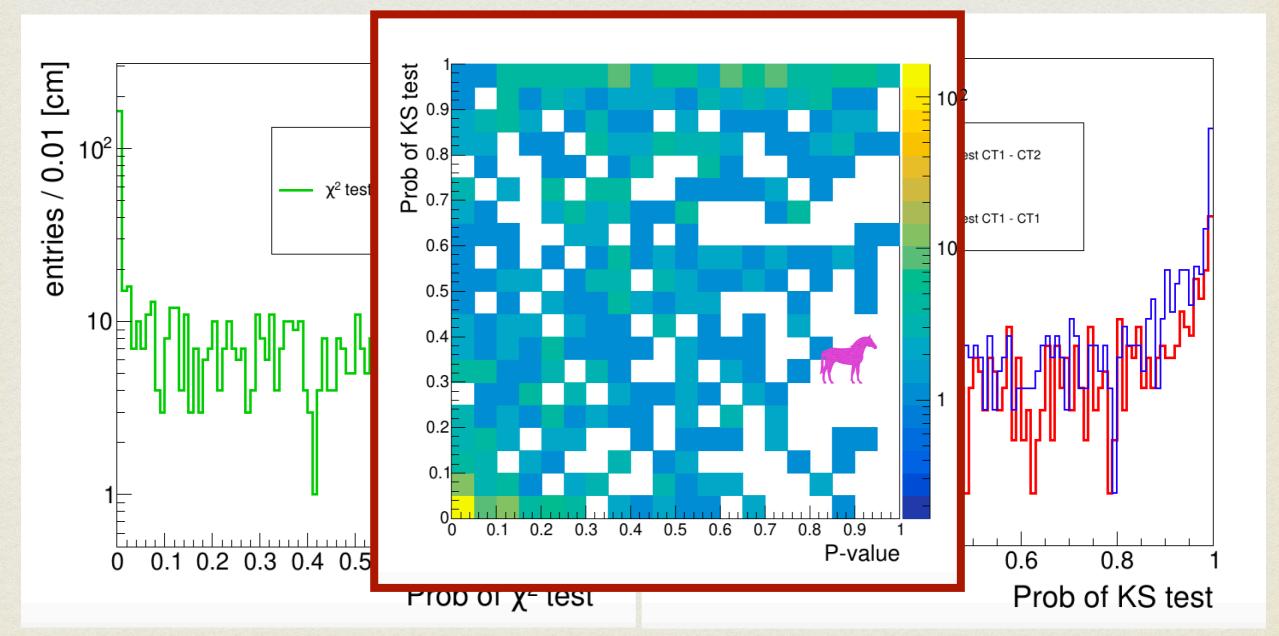
2

**B3** 



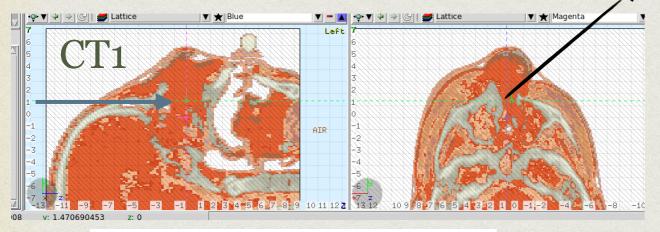


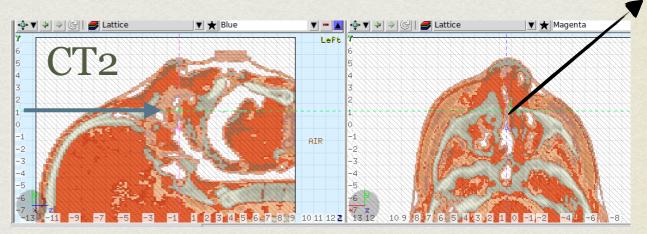
The bidimensional visualization of the  $\chi^2$  and KS probability shows a population of super PB with  $p(\chi^2)$  and p(KS) < 1%

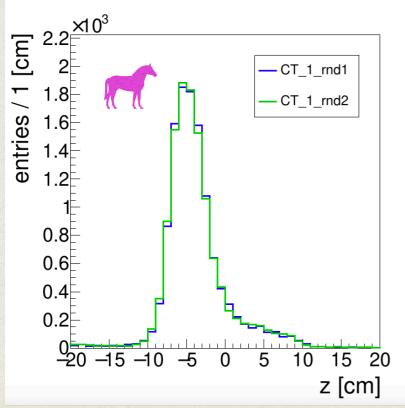


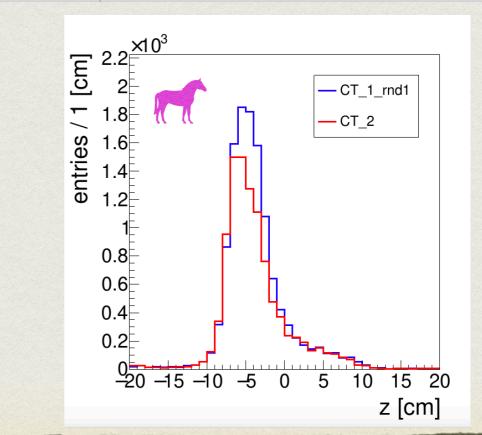
#### PB WITH KS AND x<sup>2</sup><0,01

A detailed study of individual super PBs has been done to show how the different spectra are related to the toxicity onset...

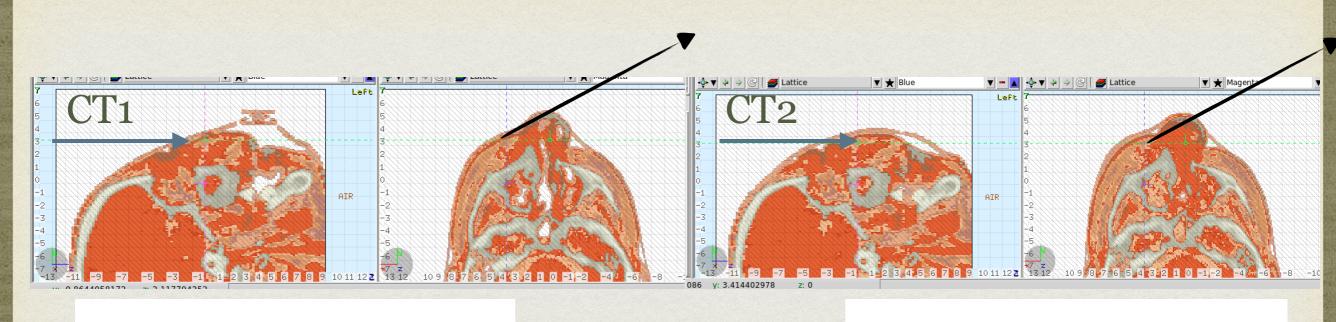


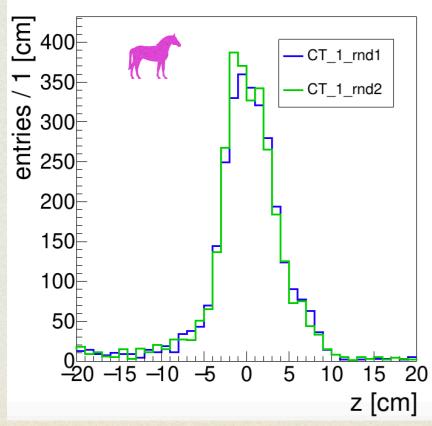


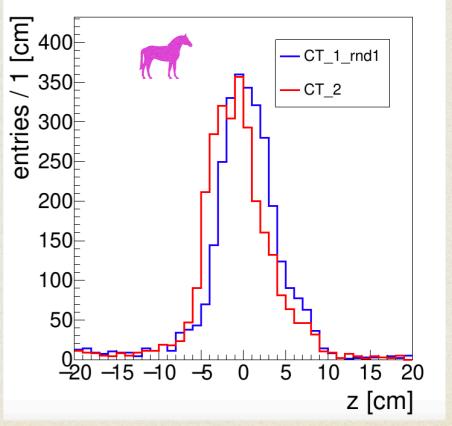




### PB WITH KS AND x<sup>2</sup><0,01

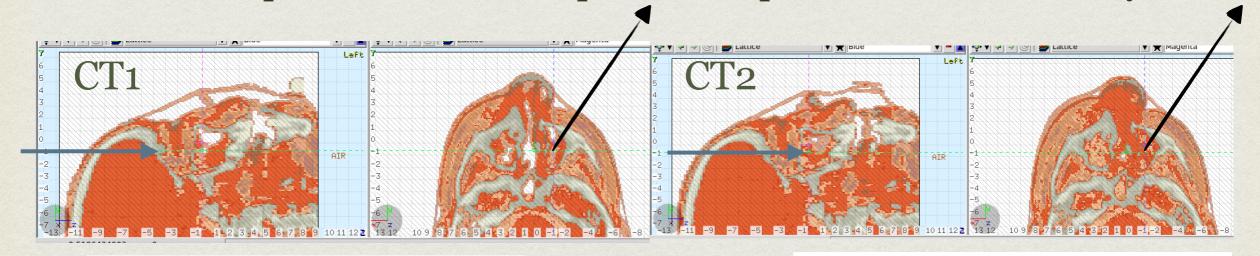


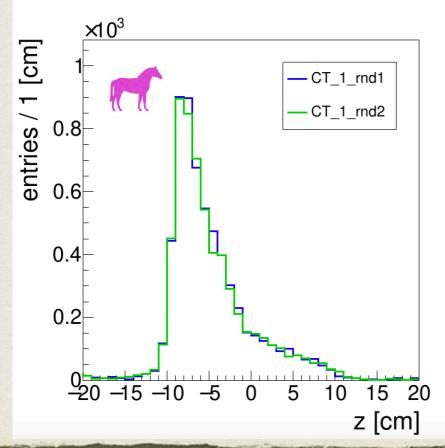


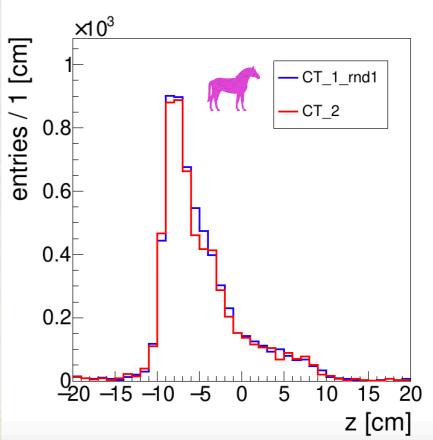


# PB WITH KS AND $\chi^2$ BETWEEN 0,6 AND 0,9

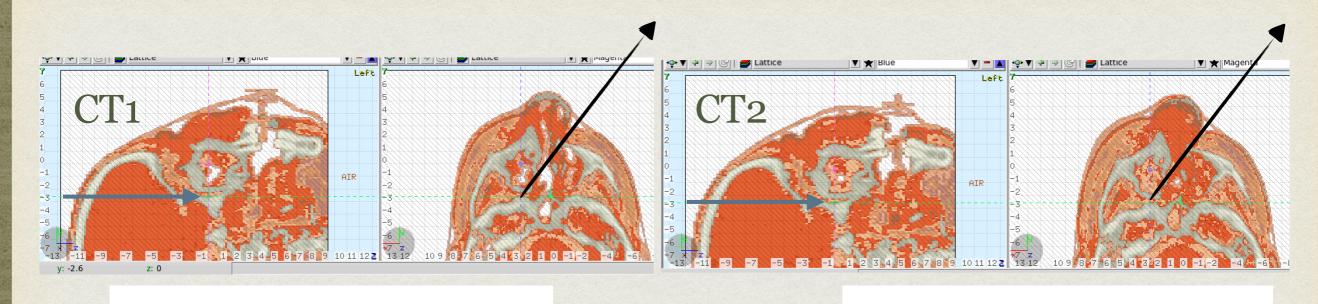
Individual super PBs that have spectra compatible have been analyzed

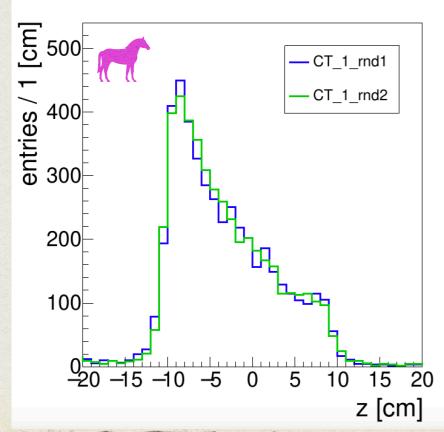


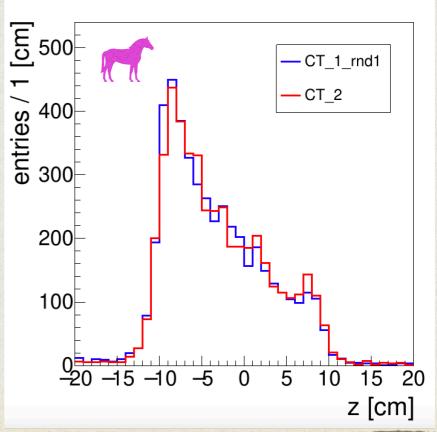




# PB WITH KS AND $\chi^2$ BETWEEN 0,6 AND 0,9





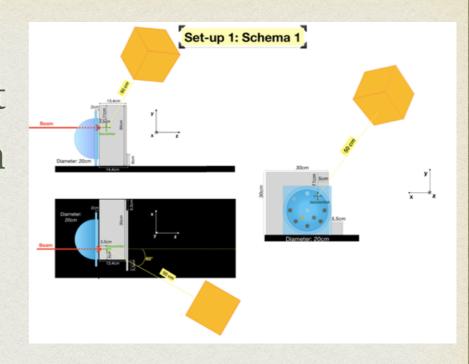


#### CONCLUSION

- The inter-fractional monitoring capability of the DP has been tested in the case of an ACC and the preliminary MC results seems to be promising
- As expected we are more sensitive in some fields due to the relative positioning of the DP wrt the target volume and the absorption inside the body
- A paper is in preparation documenting the DP capability on the basis of the FLUKA MC simulation

#### NEXT STEPS

• Perform the analysis of the data collected at CNAO @ end of 2018 with "phantom" with insets of different density



- Finalize the study of B1 of a different patient where no sensitivity to the toxicity was observed -> redo the study with the proper "CT morphing" and check the results
- Beautify the plots and finish the article preparation