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ADAPTIVE RADIOTHERAPY (A.R): RISULTATI DOSIMETRICI IN 10 PAZIENTI CON CARCINOMA DEL RINOFARINGE E DELL'OROFARINGE TRATTATI CON TECNICA VMAT

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PADOVA



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Adaptive radiotherapy

"Adaptive radiotherapy" is defined as changing the radiation treatment plan delivered to a patient during a course of radiotherapy to account for

- **Temporal changes in anatomy** (e.g. tumor shrinkage, weight loss or internal motion)
- **Changes in tumor biology/function** (e.g. hypoxia).



Adaptive RT



ELSEVIER

Seminars in
**RADIATION
ONCOLOGY**

Adaptive Radiotherapy of Head and Neck Cancer

Pierre Castadot, MD, John A. Lee, Eng, PhD, Xavier Geets, MD, PhD, and
Vincent Grégoire, MD, PhD, FRCR

Intensity-modulated radiation therapy (IMRT) in head and neck (H&N) cancer has the capability to generate steep dose gradients, leading to an improved therapeutic index. IMRT plans are typically based on a pretreatment computed tomography scan that provides a snapshot of the patient's anatomy. Nevertheless, interfractional patient variations may occur because of setup error and anatomical modifications. Therefore, the accuracy of IMRT delivery for H&N cancer may be compromised during the treatment course, potentially affecting the therapeutic index. In this framework, adaptive radiotherapy is a potential solution, which consists of "the explicit inclusion of the temporal changes in anatomy during the imaging, planning, and delivery of radiotherapy." Adaptive radiotherapy has brought an additional dimension to the management of patients with H&N cancer and has the potential to counteract the effects of positioning errors and anatomical changes. This article reviews the causes and discusses potential solutions to circumvent the discrepancies between the planned dose and the actual dose received by patients treated for H&N malignancies.

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Adaptive RT

-Buon controllo locale (no dose-escalation)

-Riduzione della dose agli OAR
(parotidi)



Adaptive RT-anatomical modifications

Table 1 Anatomical Modifications During Radiation Therapy

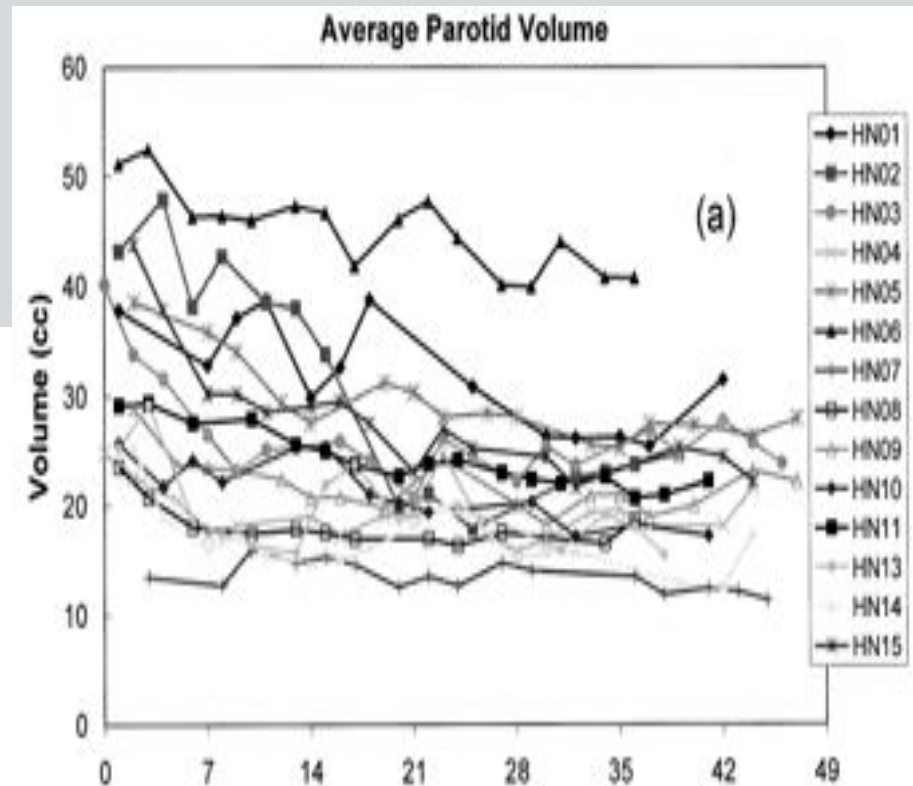
Author	No. of Patients	Per-Treatment Imaging	Image Registration	Volume Analysis	Shape and Positional Analysis
Barker et al (2004) ²⁰	14	In-room CT-on-rail 3 times/wk; no iv contrast	Rigid	Reduction of: • GTV: 1.8% per treatment day • PGs: 0.6%/treatment day	• GTV: COM displacement: 3.3 mm (asymmetric shrinkage) • PG: COM shift medially by 3.1 mm NA
Geets et al (2007) ²⁰	10	CT scan at mean doses of 14, 25, 35, and 45 Gy; iv contrast	Rigid	After a mean dose of 45 Gy: • GTV: mean decrease of 65.5% • High dose CTV: mean decrease of 50.9% • High dose PTV: mean decrease of 47.9%	NA
Han et al (2008) ⁴³	5	Daily helical MVCT	Rigid	At the end of treatment: PGs had decreased from 20.5 to 13.2 cm ³ , ie, an average decrease of 0.21 cm ³ /treatment day or 1.1%/treatment day	NA
Vasquez Osorio et al (2008) ⁵¹	10	CT scan at 46 Gy; iv contrast	Deformable	Reduction after 46 Gy: • GTV: 25 ± 15% • Homolat PG: 17 ± 7% • Heterolat PG: 5 ± 4% • Homolat SMG: 20 ± 10% • Heterolat SMG: 11 ± 7%	After 46 Gy: • Lateral and inferior regions of homolat PG: medial and posterior shift (3 mm) • Homolat SMG: medial, cranial, and posterior shift (4 mm) NA
Hansen et al (2006) ²²	13	CT scan after a mean dose of 38 Gy	Rigid	Reduction: • GTV: no change • Right PG: 15.6% • Left PG: 21.5%	NA
Robar et al (2007) ²³	15	Weekly CT scans; no iv contrast	Rigid	Reduction of superficial regions of both PGs: 4.9%/wk	Superficial regions show medial translation of: left PGs: medial shift of 0.91 ± 0.9 mm/wk right PGs: medial shift of 0.78 ± 0.13 mm/wk
Castadot et al (2008)	10	CT scan at mean doses of 14, 25, 35, and 45 Gy; iv contrast	Deformable	Reduction of • GTV: 3.2%/treatment day • GTV _N : 2.1%/treatment day • Homolateral PG: 0.9%/treatment day • Heterolat PG: 1.0%/treatment day • Low dose homolat CTV _N : 0.5%/treatment day • low dose heterolat CTV _N : 0.4%/treatment day	After 5 treatment wks: • Homolat PG: medial shift of 3.4 mm • GTV _T : lateral shift of 1.3 mm • GTV _N : medial shift of 0.9 mm • Low dose homolat CTV _N : medial shift of 1.8 mm No shift for the heterolat PG and heterolat low dose CTV _N .



Adaptive RT-anatomical modifications

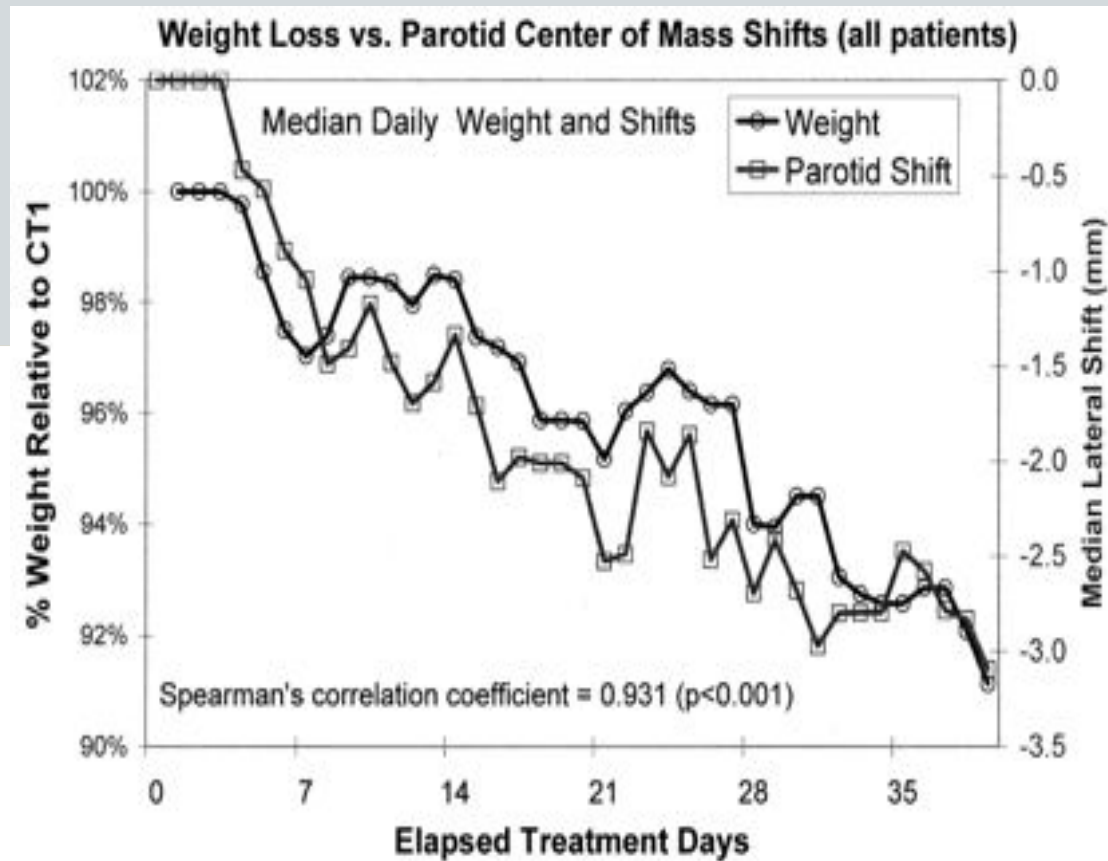
Riduzione media del volume parotideo del 28%

Cambiamenti anatomici significativi dopo 3-4 sett.



Adaptive RT-anatomical modifications

Shift mediale delle parotidi altamente correlato al calo ponderale



Adaptive RT-anatomical modifications and Dosimetric effects

Table 2 Dosimetric Effect of Anatomical Modifications During Radiation Therapy

Author	No. of Patients	Per-Treatment Imaging	Image Registration	Results	Comments
O'Daniel et al (2007) ¹⁴	11	In-room CT-on-rail scans twice/wk; no iv contrast	Deformable	Cumulative PG dose greater than planned; median dose increase: 1 Gy No impact on tumor dose coverage	If no image-guidance for daily setup error correction, cumulative PG dose greater than planned; median dose increase: 3 Gy for homolat PG and 1 Gy for heterolat PG
Hansen et al (2006) ²²	13	CT scan after a mean dose of 38 Gy	Rigid	<ul style="list-style-type: none"> decreased by 12.1, 12.2 Gy, and 7%, respectively Low dose PTV D_{30}, D_{25}, $V_{20\%}$ decreased by 12.6, 11.3 Gy, and 8.2%, respectively Right PG V_{20Gy} increased by 10.9% Mandible V_{40Gy} increased by 7.2% 	If replanning; significant improvement of: <ul style="list-style-type: none"> Low and high dose PTVs D_{30}, D_{25} and $V_{20\%}$ Spinal cord D_{max}, D_{1cc} Brainstem D_{max} Right parotid PG D_{mean}, D_{50} and V_{20Gy} Mandible D_{max} and V_{40Gy}
Robar et al (2007) ²³	15	Weekly CT scan; no iv contrast	NA	<ul style="list-style-type: none"> Left PG D_{mean} increased by $2.6 \pm 4.3\%$; V_{20Gy} increased by $3.5 \pm 5.2\%$ Right PG D_{mean} increased by $0.2 \pm 4.0\%$; V_{20Gy} increased by $0.3 \pm 4.7\%$ 	
Han et al (2008) ¹³	5	Daily helical MVCT	Rigid	PG D_{mean} increased from 0.83 to 1.42 Gy with an average increase rate of 0.17 Gy/treatment day corresponding to an average increase of 2.2%/treatment day	Strong correlation between the volume and the median parotid dose during the treatment (correlation coefficient, -0.95)
Lee et al (2008) ²⁴	10	Daily helical MVCT	Deformable	<ul style="list-style-type: none"> PG daily D_{mean} differed from the planned dose by an average of 15% PG cumulative D_{mean}: planned: 29.7 Gy actual: 32.7 Gy (110% of planned dose) 	<ul style="list-style-type: none"> Changes in the distance between the COMs of the left and right PGs correlated strongly with the mean parotid dose changes ($R^2 = 0.89$) Correlation between the relative weight loss and higher parotid mean doses ($R^2 = 0.58$)
Castadot et al (2009)	10	CT scan at mean doses of 14, 25, 35, and 45 Gy; iv contrast	Deformable	PGs D_{mean} : planned: 17.9 Gy, actual 18.7 Gy SMGs D_{mean} : planned 51.9 Gy actual: 52.8 Gy OC D_{mean} : planned 26.0 Gy, actual 26.7 Gy SC D_2 : planned 40.1 Gy, actual: 41.0 Gy Skin V_{40} : planned 17.2 Gy, actual 18.3 Gy No difference in PTV or CTV coverage	



Adaptive RT

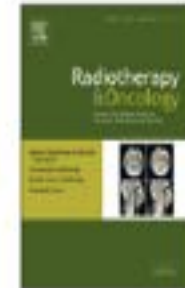
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Adaptive RT in head and neck cancer

Adaptive functional image-guided IMRT in pharyngo-laryngeal squamous cell carcinoma: Is the gain in dose distribution worth the effort?

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Adaptive RT

-Clinical benefit of adaptive strategy will have to be evaluated both in terms of patients outcome and cost-effectiveness

-In the mean-time, owing to the complexity of the procedure, **ART should not be used in routine clinical base**



Adaptive RT-Padova



MATERIALI E METODI

-10 pazienti tra 2013-2014

-III-IV stadio rinofaringe e orofaringe

ID pz	Istologia	Sesso	Eta	Stadio	CT concomitante	Kg.persi	N* TC
1	SCC	M	70	T3N2c	NO	16	5
2	NPC	M	70	T3N1	SI	7	6
3	SCC	F	67	T3N0	SI	7	5
4	SCC	M	58	T2N2b	SI	14	5
5	NPC	M	36	T2N1	SI	6	6
6	NPC	M	63	T2N1	SI	5	6
7	NPC	M	68	T3N2	SI	9	6
8	NPC	F	69	T1N2	SI	4	6
9	NPC	F	59	T2N2	SI	6	6
10	NPC	M	40	T2N2	SI	10	6

MATERIALI E METODI

-VMAT a 2 archi(360 gradi)

Varian Unique

-Pianificazione con **Eclipse** TPS

-5mm di margine per il PTV

66Gy-60-54/30f SIB per Orofaringe

69.9Gy-60-54/33f SIB per Rinofaringe

-TC settimanale di **controllo**

-Registrazione elastica con la TC di pianificazione
(**Raystation** TPS)

Workflow

Simulazione

Pianificazione
P1

Trattamento

• TC settimanale di controllo

Registrazione
elastica

• Calcolo del piano P1
per le restanti sedute
sulla TC di controllo
deformata sulla TC di
pianificazione

Cumulative
dose

• Ripianificazione
P2 o meno

Ripresa del
trattamento

MATERIALI E METODI

-9 pazienti sono andati incontro ad ART1

-Tempo medio tra TC e ART1 :1.8 giorni

-Media 1[^] ripianificazione: 16[^]frazione

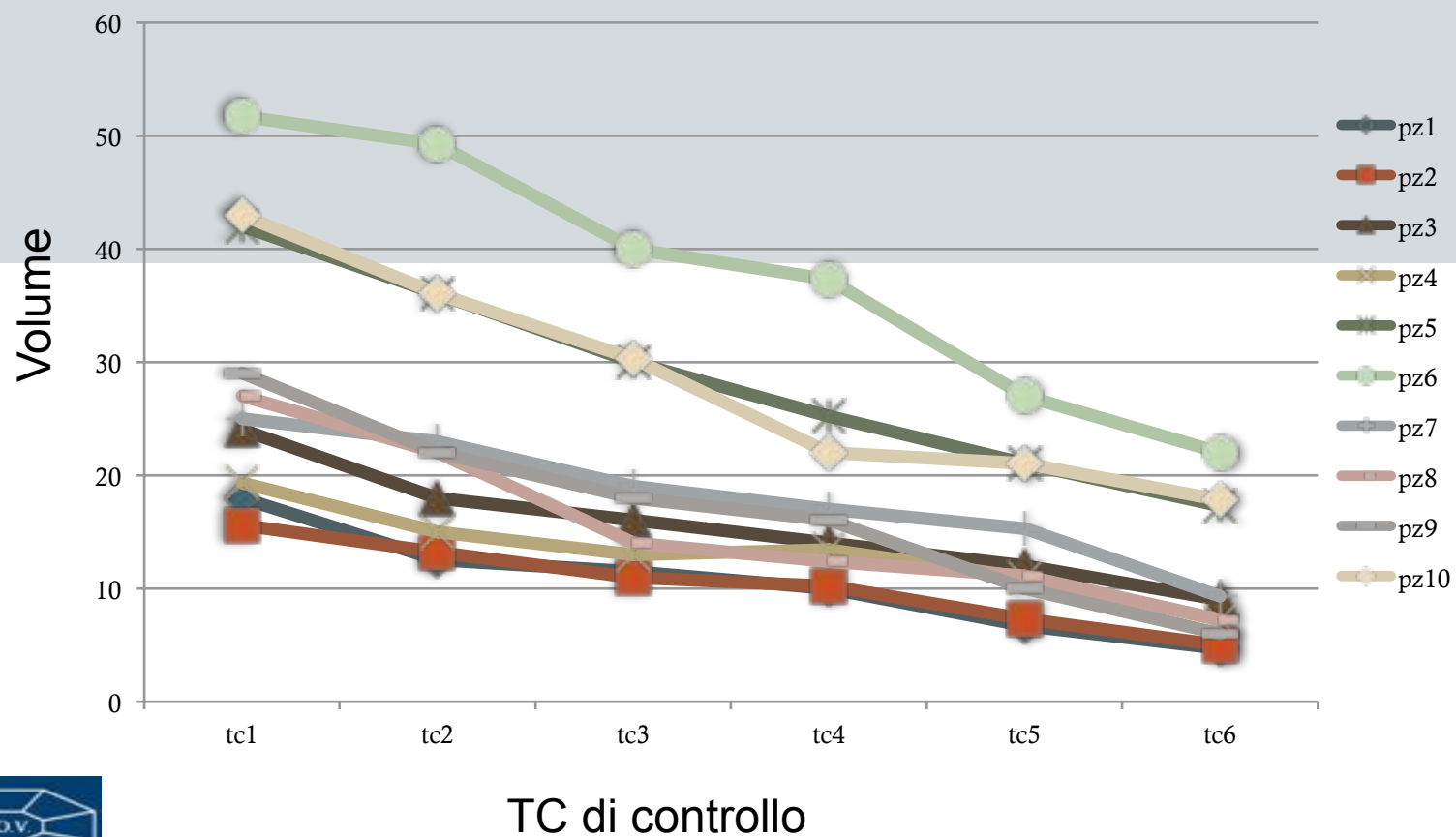
-2 pazienti(orofaringe) ART2

-Media 2[^]ripianificazione:22[^]frazione

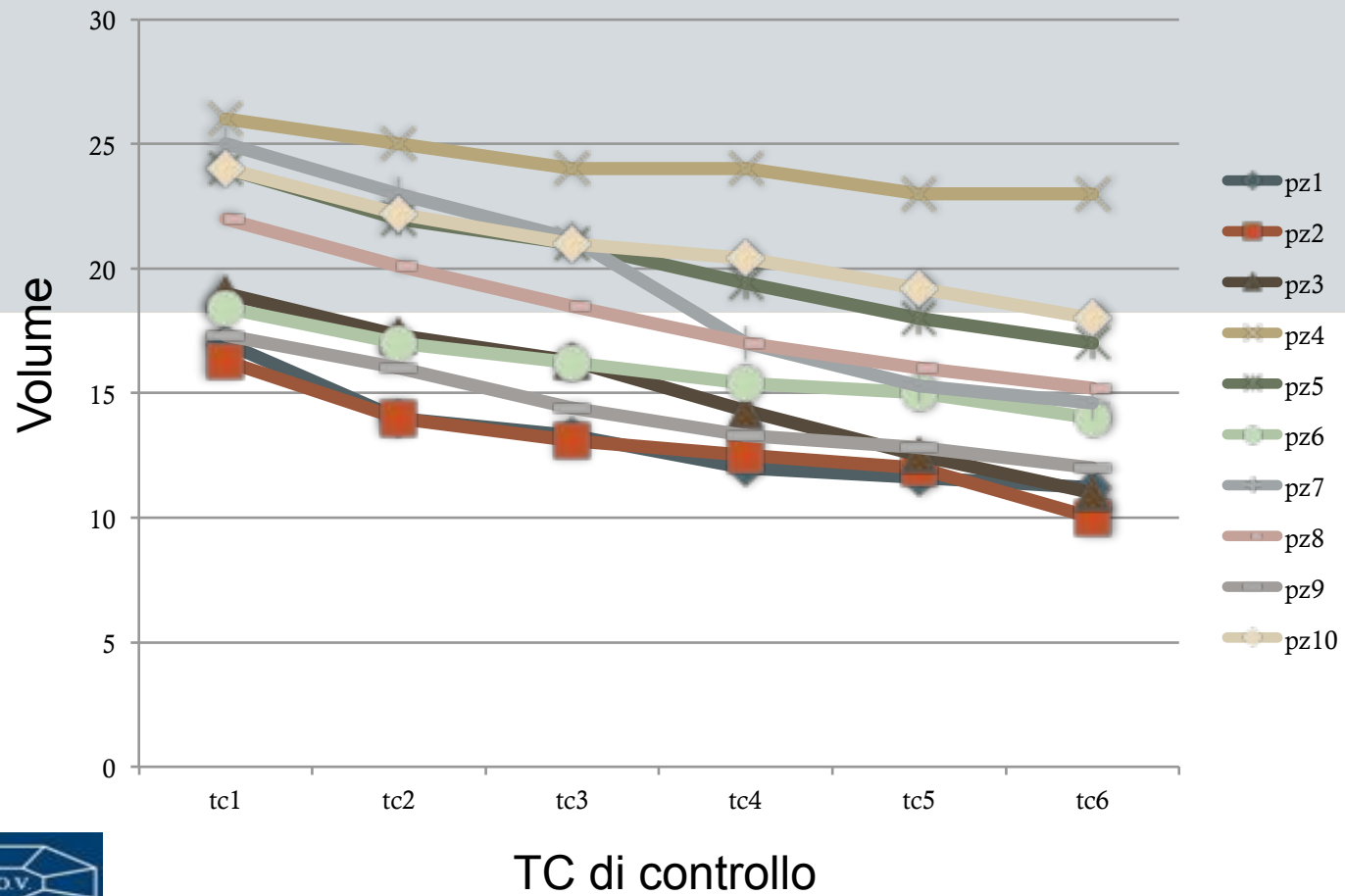
RISULTATI

- Volume del GTV diminuisce del ordine del 35% durante le prime 3 settimane di trattamento
- Volume medio parotideo diminuito dell 11%
- ART 1 riduce la dose media alle parotidi di 1,1 Gy e al tronco di 0,8 Gy
- ART 2 nei casi Ca orofaringe riduce la dose media alla parotide vicina al GTV di circa 3 Gy

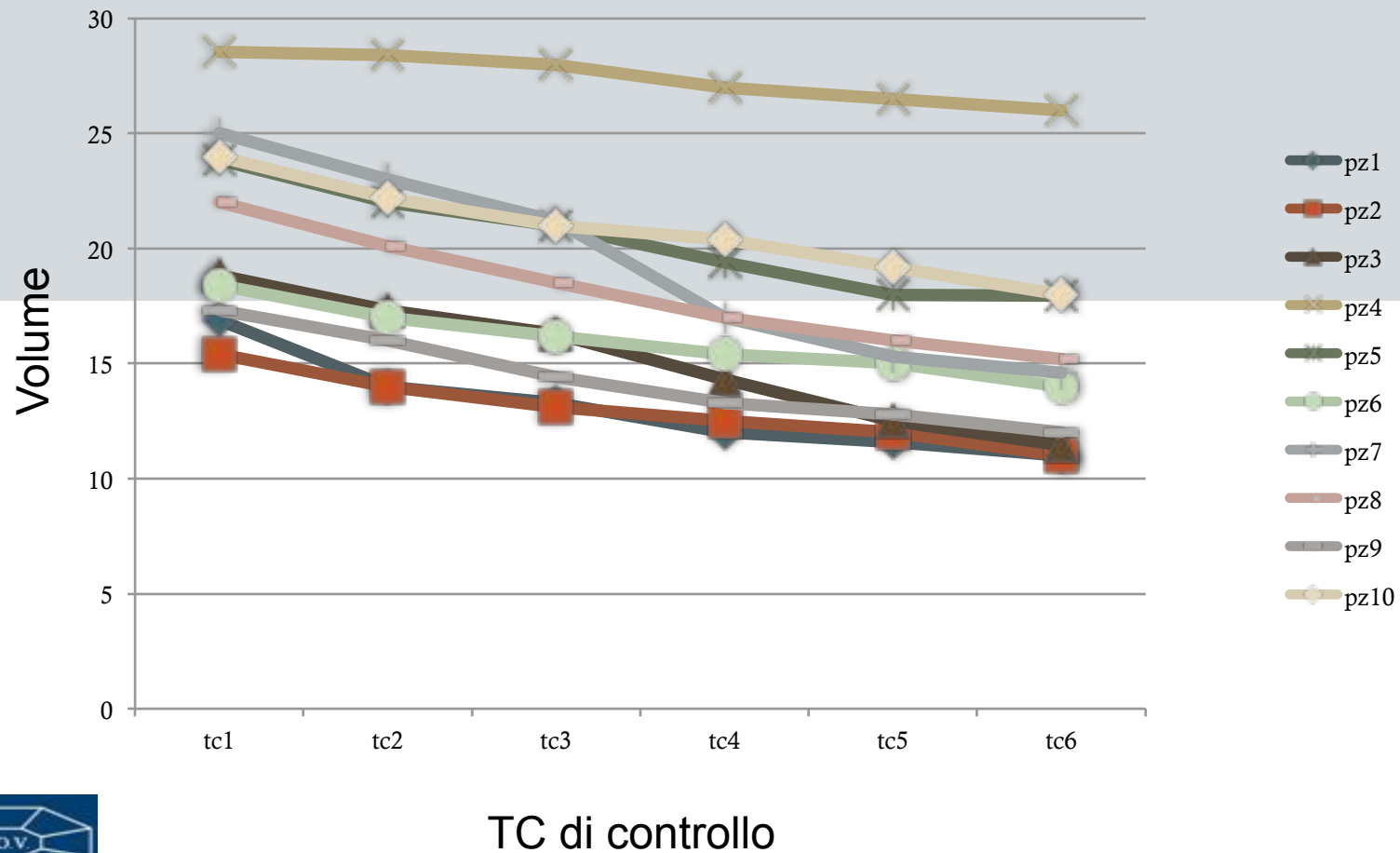
Modifica del GTV



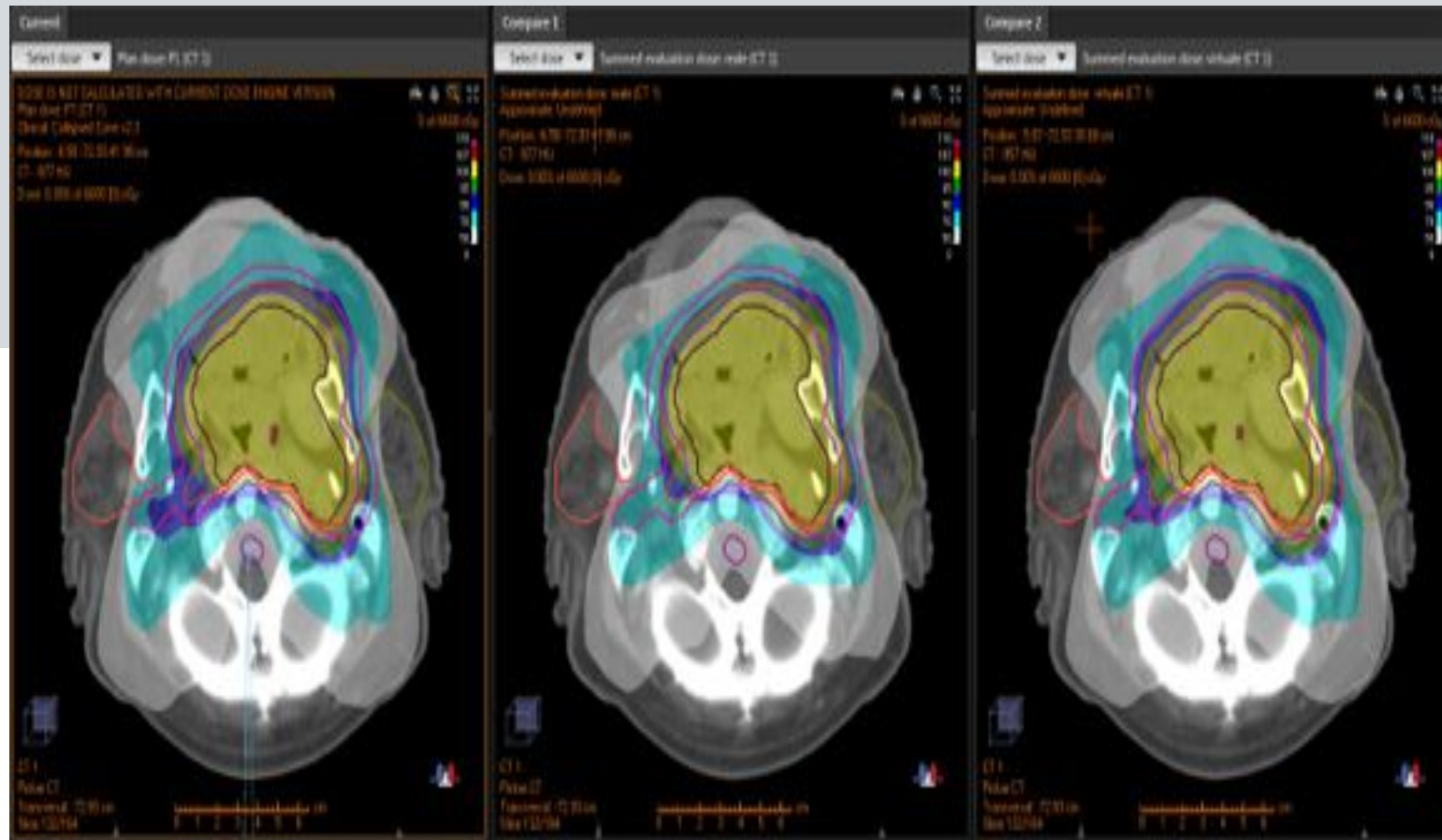
Volume parotideo sx



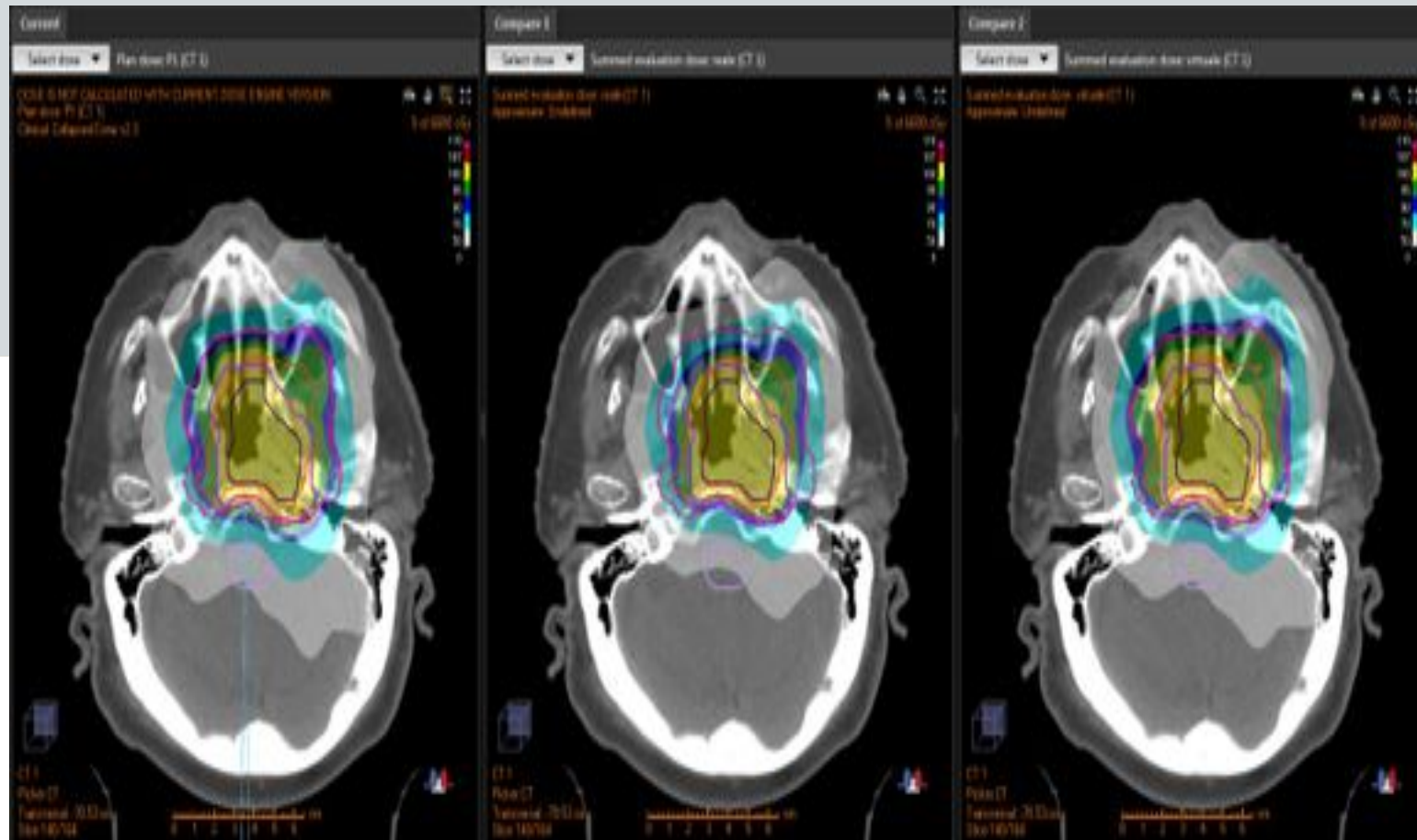
Volume parotideo sx



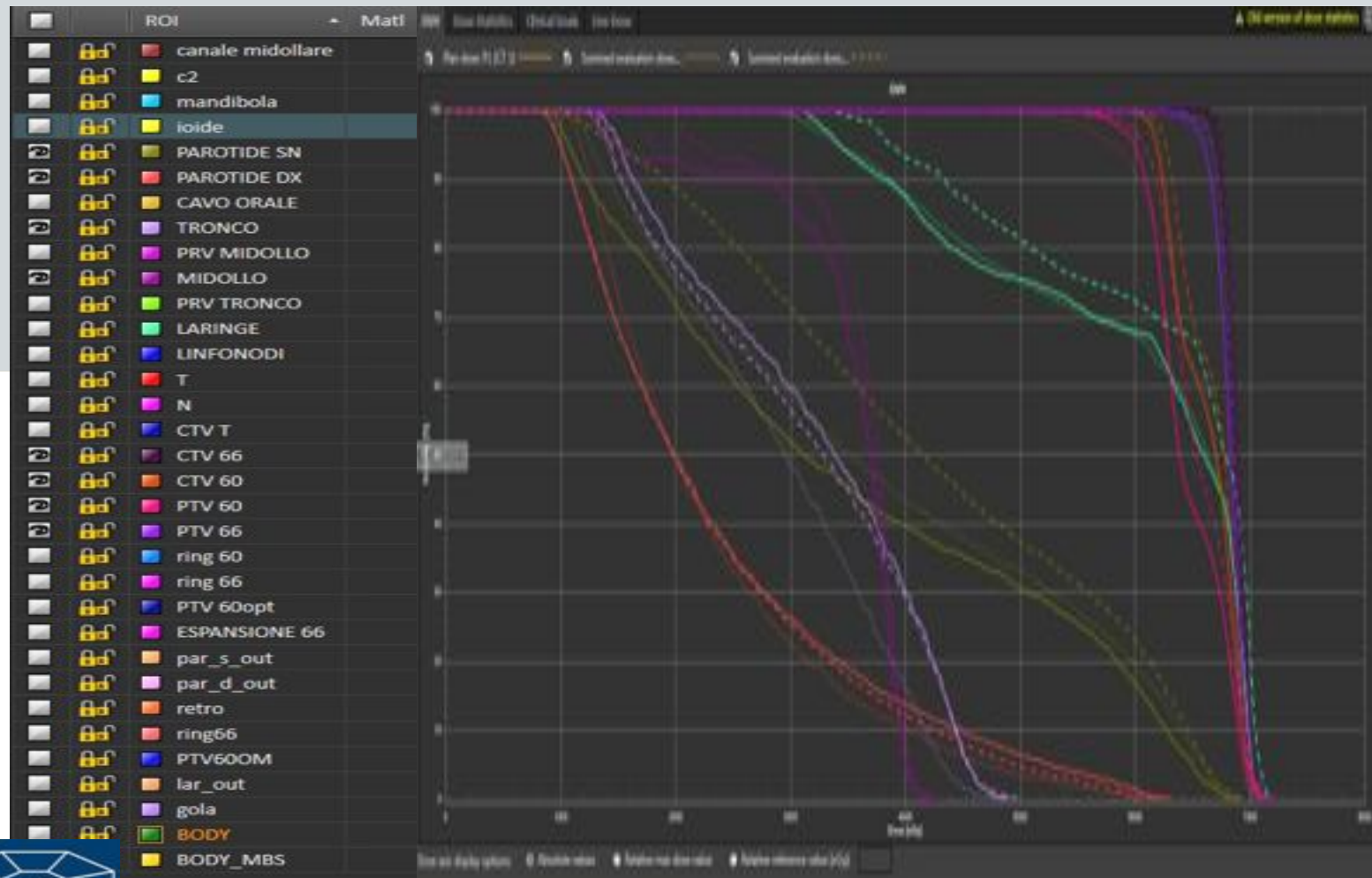
Risultati-caso 1



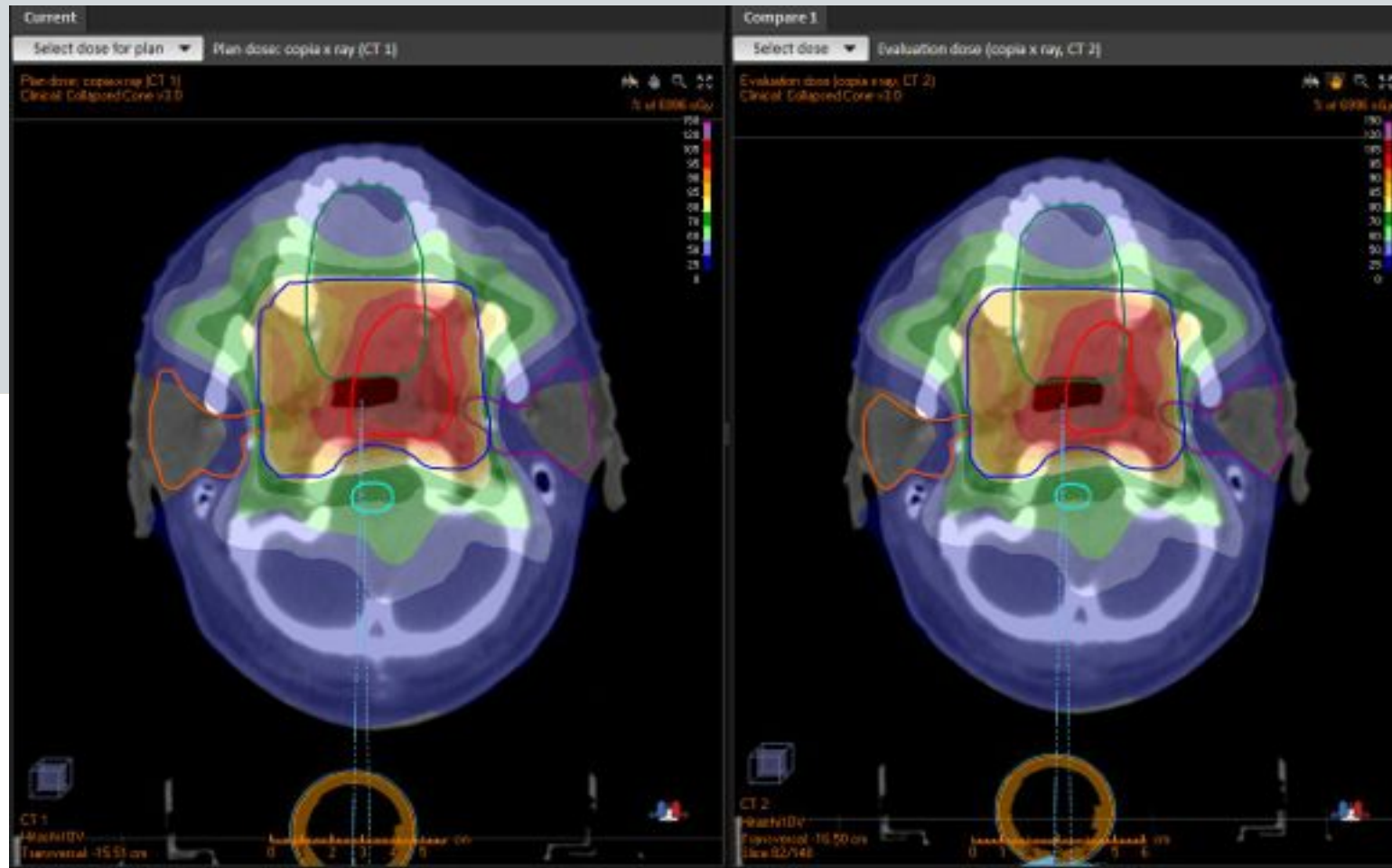
Risultati-caso 1



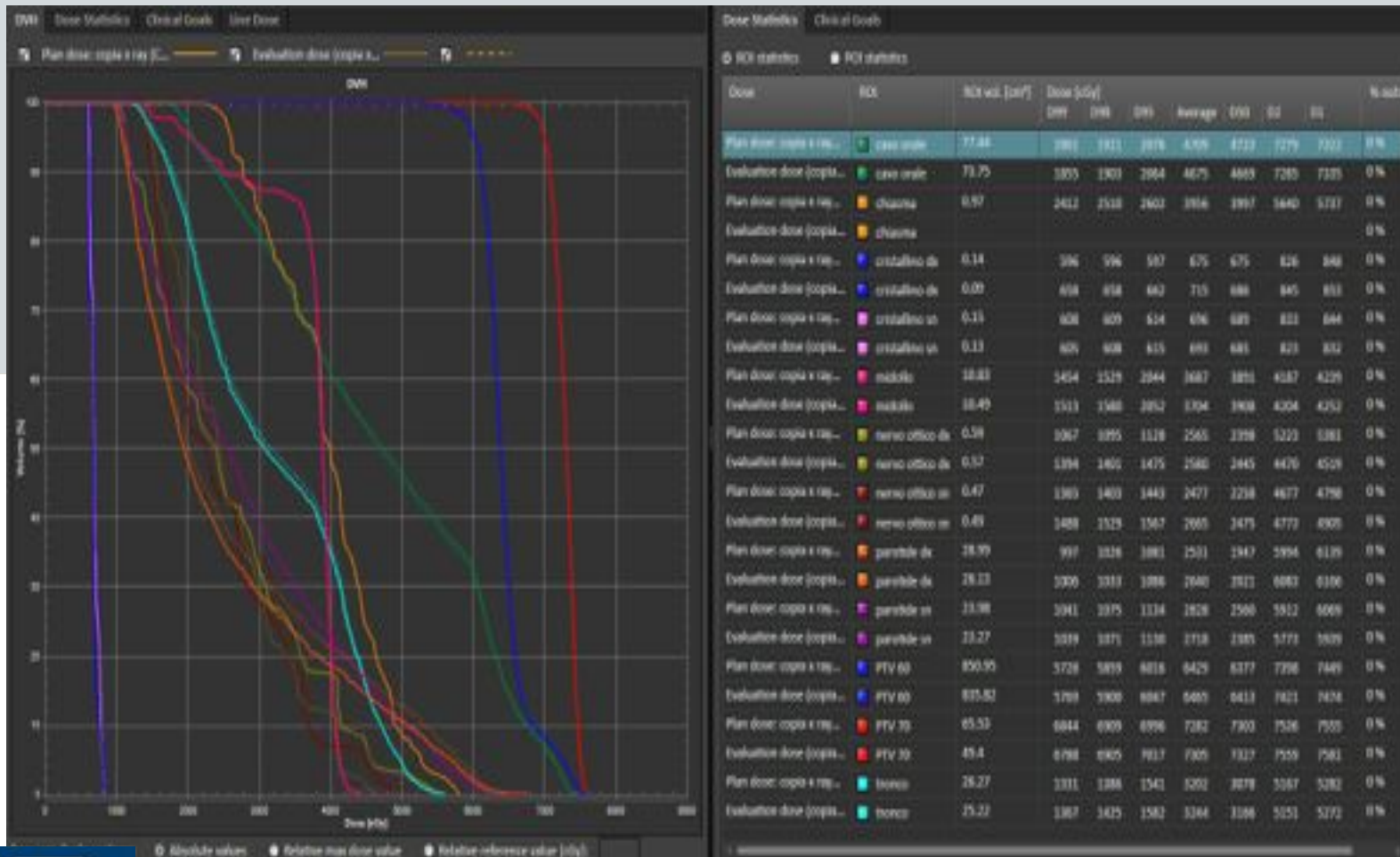
Risultati-caso1



Risultati-caso 2



Risultati-caso 2



CONCLUSIONI

- L' ART e' fattibile
- Grande impegno di tempo e risorse umane
- Una migliore conoscenza degli algoritmi di registrazione elastica e dei loro limiti è necessaria prima che 'A.R. possa essere introdotta con sicurezza nella pratica clinica giornaliera
- Nel nostro studio, modesto beneficio nella ripianificazione adaptive in pazienti non selezionati.
- L'impatto clinico di questa tecnica deve essere ancora confermato da futuri trials.



GRAZIE !!!