School on Medical Physics for Radiation Therapy: Dosimetry, Treatment Planning and Delivery for Advanced Applications

11 - 22 September 2023 An ICTP Meeting Trieste, Italy

Further information: http://indico.ictp.it/event/10205/ smr3871@ictp.it

**(CTP** 

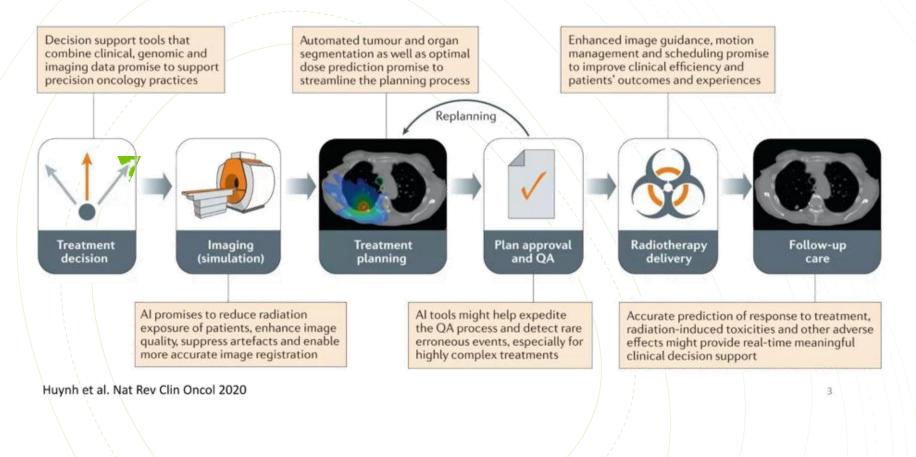
#### Artificial intelligence in Radiotherapy

Michele Avanzo

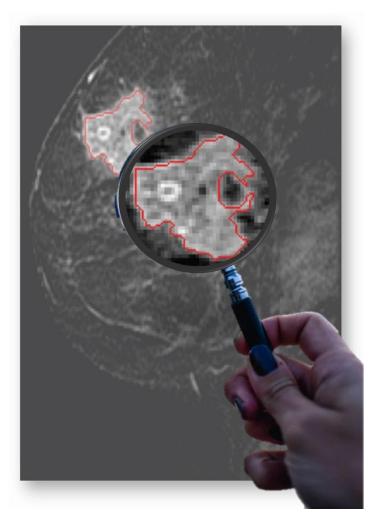
Centro di Riferimento Oncologico di Aviano IRCCS, Aviano (PN), Italy



#### AI in Radiation Oncology



# Treatment decision



#### Treatment decision

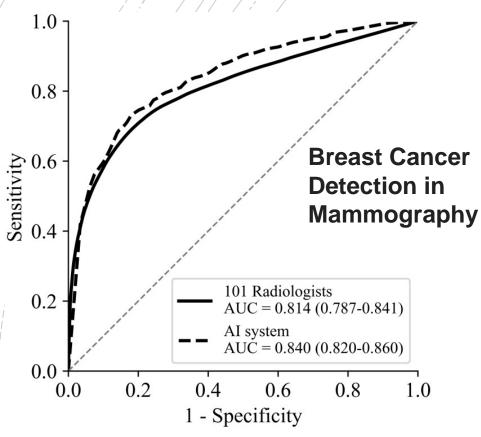


Figure 1. Receiver operating characteristic curve comparison between the reader-averaged radiologists and the artificial intelligence (AI) system in terms of area under the curve (AUC). Parentheses show the 95% confidence interval of the AUC.

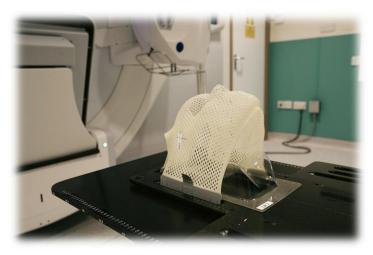
#### ChatGPT in glioma adjuvant therapy decision making: ready to assume the role of a doctor in the tumour board?

Julien Haemmerli <sup>1</sup>, <sup>1</sup> Lukas Sveikata,<sup>2,3,4</sup> Aria Nouri,<sup>1</sup> Adrien May,<sup>1</sup> Kristof Egervari,<sup>5</sup> Christian Freyschlag,<sup>6</sup> Johannes A Lobrinus,<sup>5</sup> Denis Migliorini,<sup>7</sup> Shahan Momjian,<sup>1</sup> Nicolae Sanda,<sup>2</sup> Karl Schaller,<sup>1</sup> Sebastien Tran,<sup>8</sup> Jacky Yeung,<sup>9</sup> Philippe Bijlenga<sup>1</sup>

Rodriguez-Ruiz, JNCI J Natl Cancer Inst, 2019

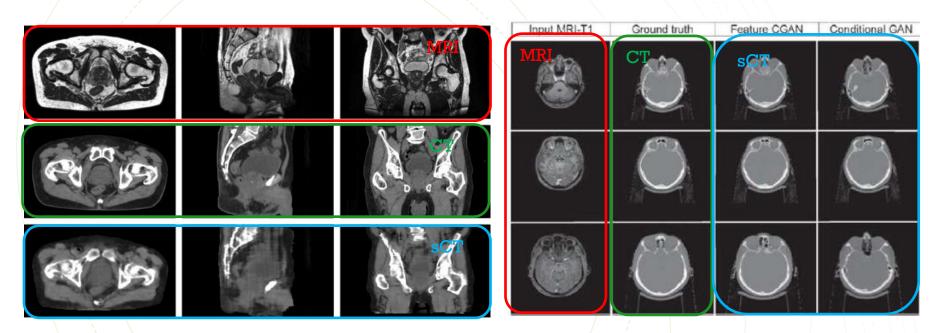






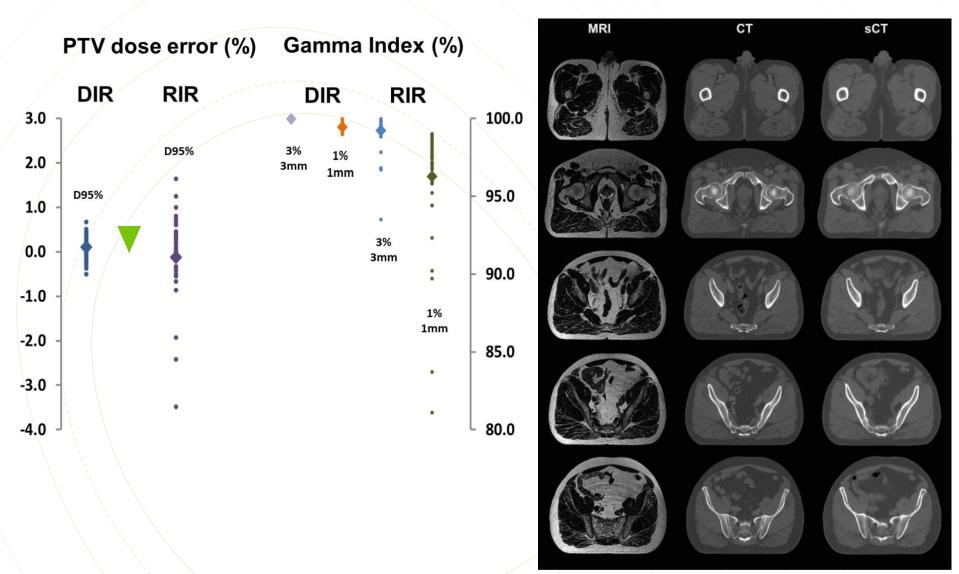
#### Conversion among imaging modalities

GAN and autoencoders



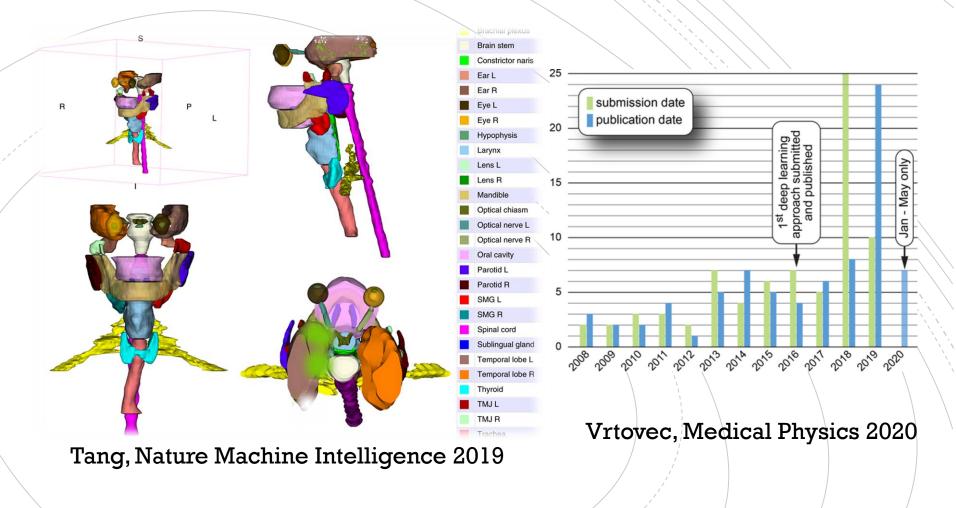
Cusumano et al. Radiotherapy and Oncology Volume 153, December 2020, Pages 205-212 Fard et al., Computers in Biology and Medicine , Vol. 146, July 2022, 105556

### Synthetic-CT generation for MR-only RT



Bird, Radiotherapy and Oncology, 2021

#### Autocontouring: head and neck



## Autocontouring: head and neck

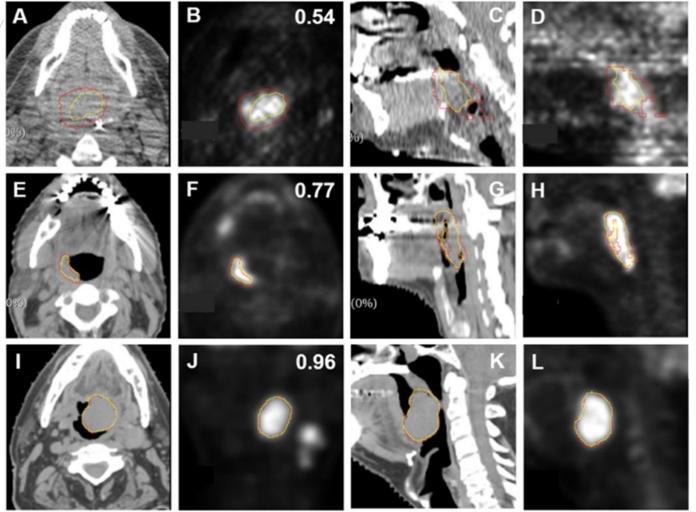
Gold standard	Model prediction	Contour comparison
		Brachial plexus Trachea Spinal cord Thyroid
		Gold standard Model
		Constrictor naris Larynx Mandible Oral cavity SMG L SMG R Spinal cord
8 S	y s	Gold standard Model
		Layrx Mardible Oral cavity Parotid L Parotid L Parotid I Spinal cord Sublingual gland
* ÷	1	Gold standard Model
		Brain stem Ear L Ear L Ear L ThU L ThU R
		Gold standard Model

OAR	MAS	AnatomyNet	U <sub>a</sub> -Net	Human
Brachial plexus	30.38±15.63	50.41±8.08	56.15 ± 10.83	33.03±7.83
Brain stem	82.25±7.47	82.63±4.57	86.25 ± 3.86	83.25±4.63
Constrictor naris	66.38±8.21	73.68±7.56	75.46 ± 6.13	62.34±8.63
Ear L	$70.38 \pm 14.94$	$76.68 \pm 5.00$	77.28 ± 4.25	43.57±12.63
Ear R	$70.03 \pm 15.57$	78.77 <u>+</u> 5.77	$78.64 \pm 6.35$	39.71±10.81
Eye L	85.96±10.99	88.41 ± 3.10	92.51 ± 2.00	90.71±2.11
Eye R	82.68±17.38	89.25±3.38	92.49 <u>+</u> 2.34	91.51±1.79
Hypophysis	$43.54 \pm 18.45$	56.18±10.01	63.86 <u>+</u> 8.73	59.26±14.77
Larynx	82.60 ± 8.19	83.06±7.98	89.25 <u>+</u> 3.26	$68.60 \pm 6.59$
Lens L	$46.25 \pm 24.29$	77.25±7.92	81.90 ± 6.88	64.27±10.06
Lens R	$45.53 \pm 23.94$	78.06±7.51	83.04 ± 5.90	71.79±9.59
Mandible	83.95±11.48	91.97±1.71	93.12 ± 1.41	90.97±1.46
Optic chiasm	42.08±17.52	60.55±11.16	64.21±16.39	28.61±14.40
Optic nerve L	59.49±14.61	$72.55 \pm 6.55$	75.73±7.26	$65.10 \pm 8.44$
Optic nerve R	59.08±16.53	72.95±7.90	76.06 ± 6.49	66.14±7.29
Oral cavity	86.10 ± 9.11	87.69±5.67	90.77 ± 2.32	79.30±3.59
Parotid L	72.52±15.57	82.28±6.71	84.86 ± 4.22	$78.46 \pm 4.90$
Parotid R	71.20±17.55	82.20 ± 7.26	84.93 <u>+</u> 3.99	78.88±4.41
SMG L	60.89±12.11	75.47±8.93	80.71 ± 7.32	77.73±6.25
SMG R	63.70±15.80	74.82±14.69	82.54 <u>+</u> 7.47	74.10 ± 16.92
Spinal cord	77.42±16.70	$80.32 \pm 6.48$	85.64 ± 5.90	84.59±6.62
Sublingual gland	21.52 ± 16.34	39.94 ± 21.02	45.99 ± 18.84	35.16±23.87
Temporal lobe L	80.05±7.28	81.76±5.33	84.78 ± 2.62	82.41±5.01
Temporal lobe R	78.26±7.40	72.97±14.60	84.13 ± 3.34	80.90±7.49
Thyroid	63.68±19.65	71.82±11.40	85.62 <u>+</u> 4.63	82.42±6.16
TMJ L	61.26±19.86	86.65±3.34	87.96 ± 3.12	84.67±5.09
TMJ R	$63.45 \pm 20.48$	85.73±3.69	86.86±3.60	81.98±8.59
Trachea	65.86±18.75	79.34 ± 7.75	$81.29 \pm 4.84$	91.05±1.69
Average	64.87	76.19	80.43	70.38

Values are given in units of %. L, left; R, right; SMG, submandibular gland; TMJ, temporomandibular joint; MAS, multi-atlas segmentation. Bold numbers represen oncologist modified his/her previous delineation by referencing the corresponding MRI images.

#### Tang, Nature Machine Intelligence 2

#### Autocontouring: bhead and neck



256 and 512 ResUnet mod

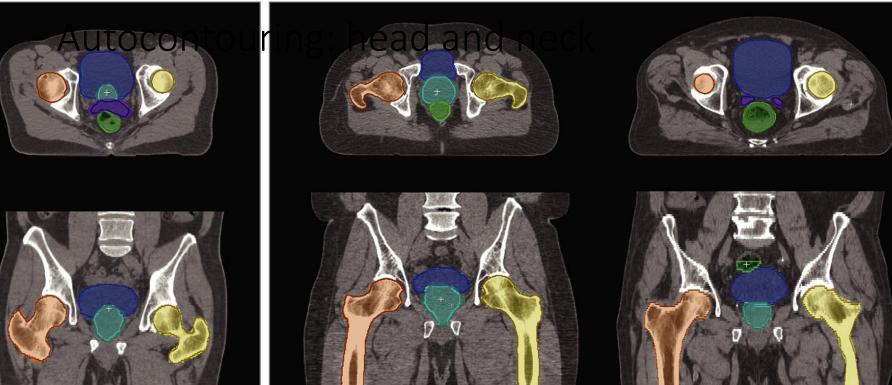
Model	DSC	
256 (mean)	$0.771\pm0.039$	
256 (median)	$0.829\pm0.024$	
512 (mean)	$0.768\pm0.041$	
512 (median)	$0.828\pm0.024$	

1

Naser, Head Neck Tumor Segm Chall (2021)





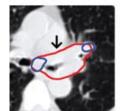


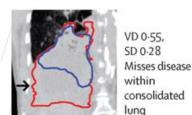
## Autocontouring: pelvis

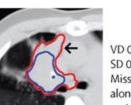
Oktay, JAMA Netw Open. 2020

#### A Model failure modes

#### **Under-segmentation**







VD 0-57, SD 0-23 Misses disease along pericardium

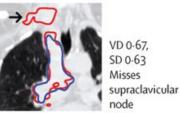
VD 0-30,

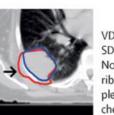
SD 0.30

No disease

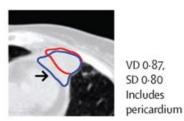
bronchus

tracking along

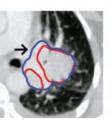




VD 0-83, SD 0.59 Not crossing ribs along pleura and chest wall



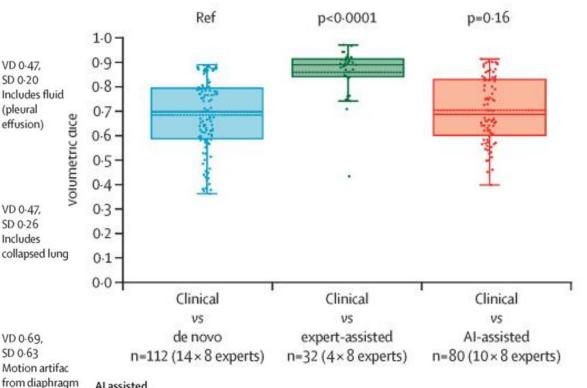
Over-segmentation



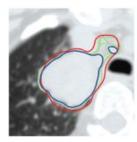
VD 0.92, SD 0-85 Includes vasculature



#### A Quantitative scoring



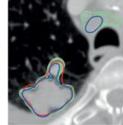
Al assisted



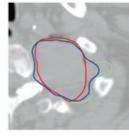
VD 0.59, SD 0.29

- Clinical — AI

— Eight experts



VD 0.76, SD 0.63

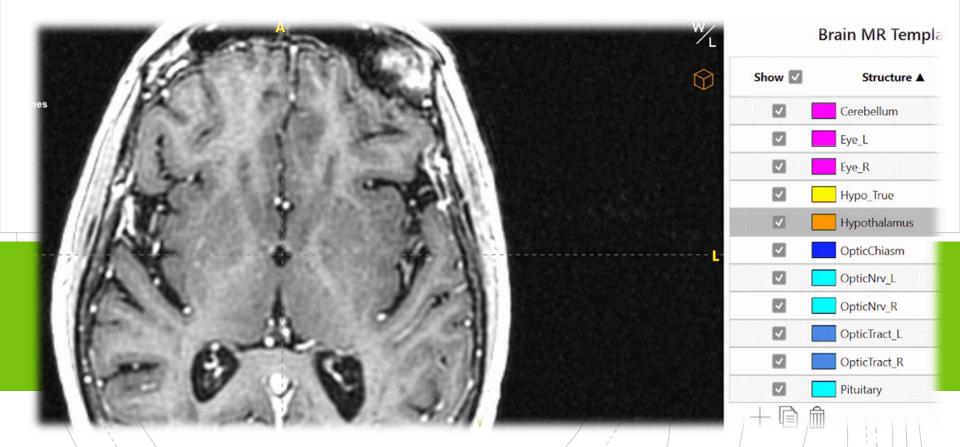


3

VD 0.91, SD 0.75

Autocontouring: failure modes

#### Autocontouring: brain



https://blog.radformation.com/autocontour-adds-to-the-list-of-available-structures

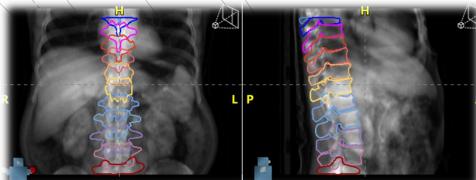
#### Autocontouring: abdomen



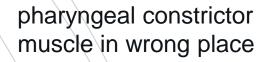
https://blog.radformation.com/autocontour-adds-to-the-list-of-available-structures

#### Autocontouring: MRI, DRR





https://blog.radformation.com/autocontour-adds-to-the-list-of-available-structures



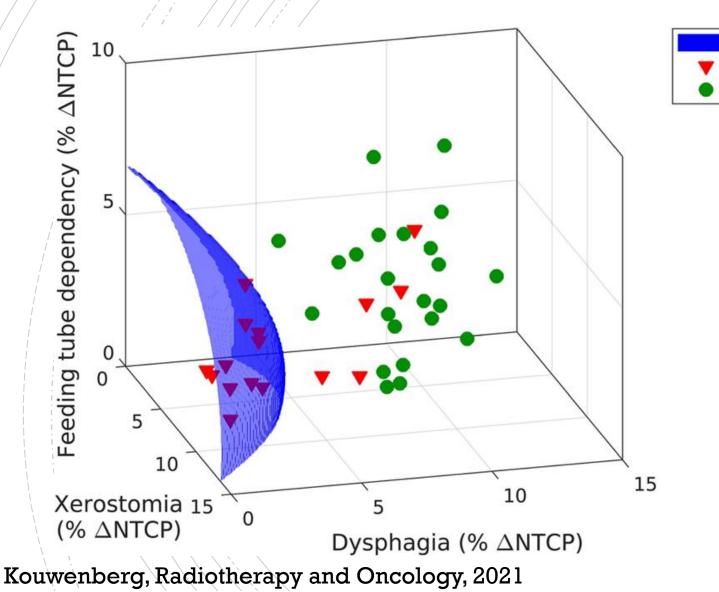
Michael Gensheimer

#### Patient selection for proton therapy

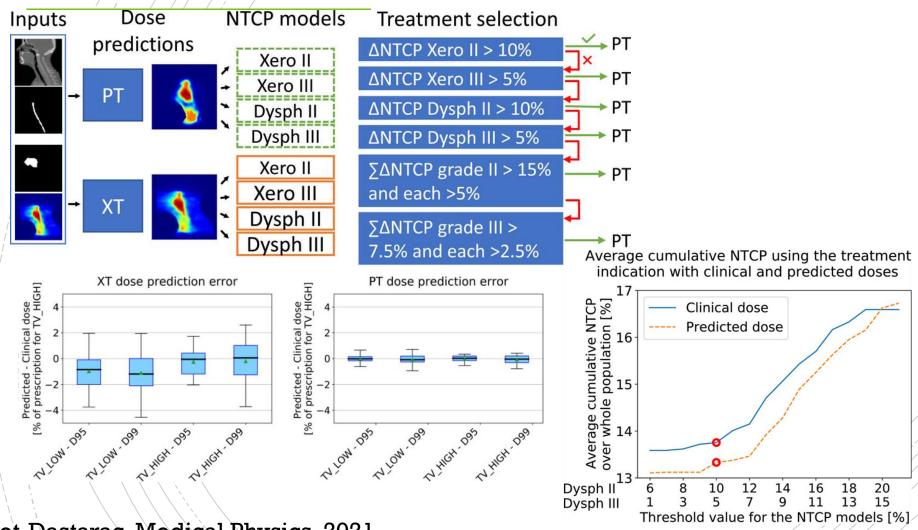
Decision boundary

not eligible for PT

eligible for PT

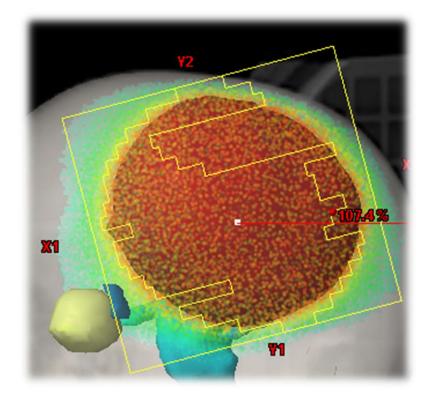


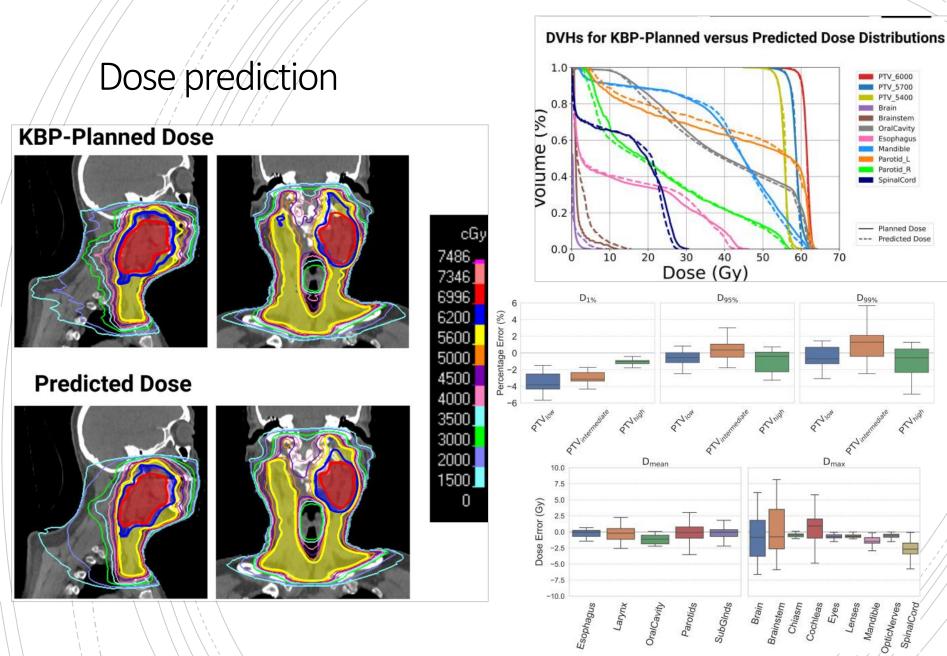
#### Patient selection for proton therapy



Huet-Dastarac, Medical Physics, 2021

#### Radiotherapy treatment planning





Gronberg, Practical Rad. Onc. 2023 

PTV\_6000 PTV\_5700 PTV 5400

Brainstem

OralCavity

Esophagus Mandible Parotid L Parotid R

SpinalCord

 Planned Dose --- Predicted Dose

70

25/10

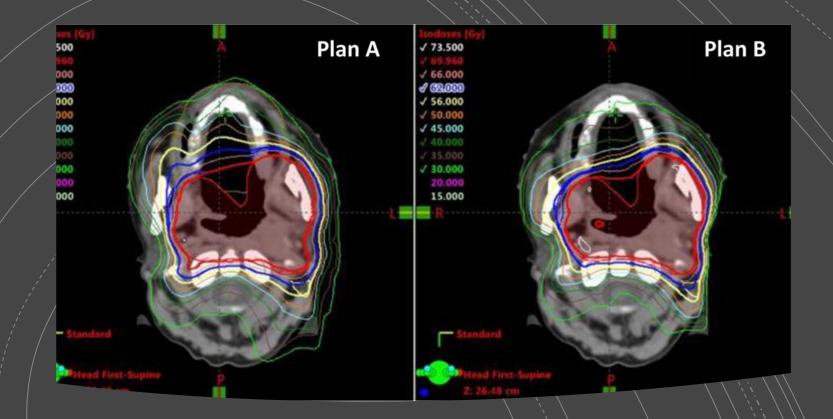
Dmax

Eyes Lenses

D99%

Mandible **OpticNerves** SpinalCord

Brain

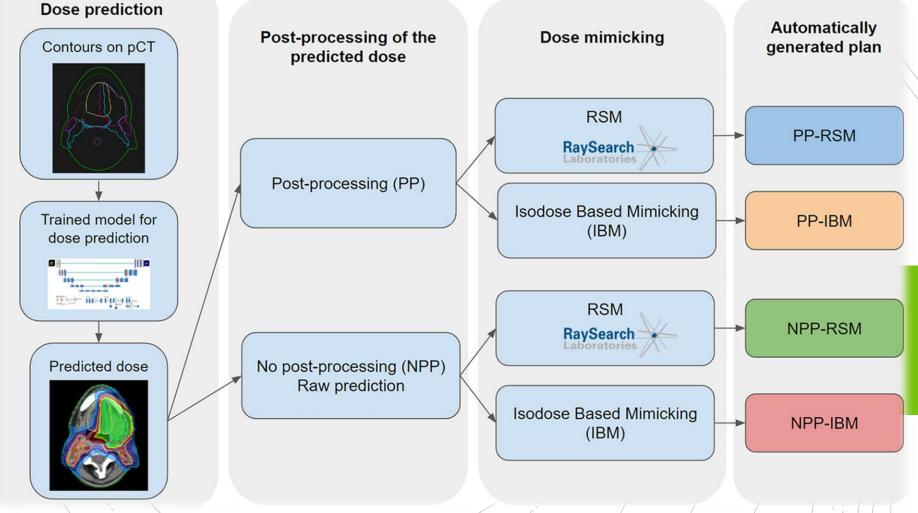


#### Automated planning

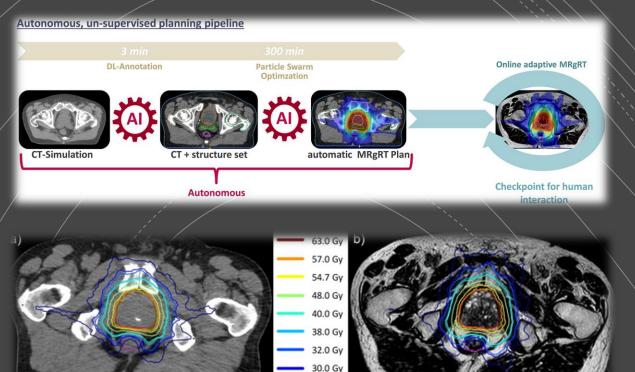
Figure clinical plan vs autoplan. Both were judged to be acceptable plans, with preference for the autoplan based on reduced dose to the oral cavity and posterior neck.

Olanrewaju, Pract Radiat Oncol. 2021

#### Dose prediction/autoplanning



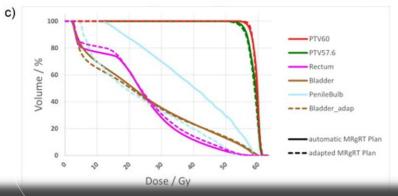
Borderias-Villarroel, Medical Physics 2023



### Fully automated RT workflow

 Autonomous plan generation, including OAR segmentation, target generation and optimization

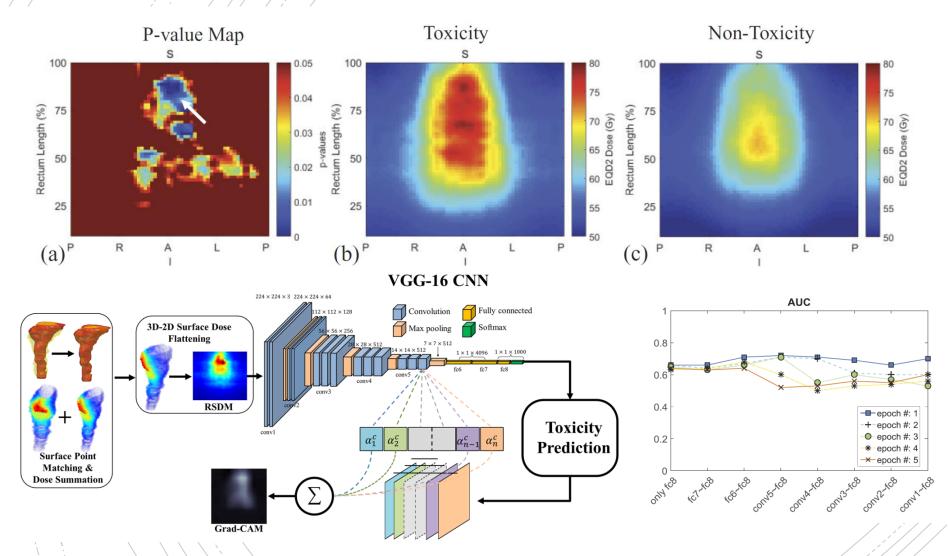
 First checkpoint for human interaction at the time of clinical online adaptation at the MR-Linac.



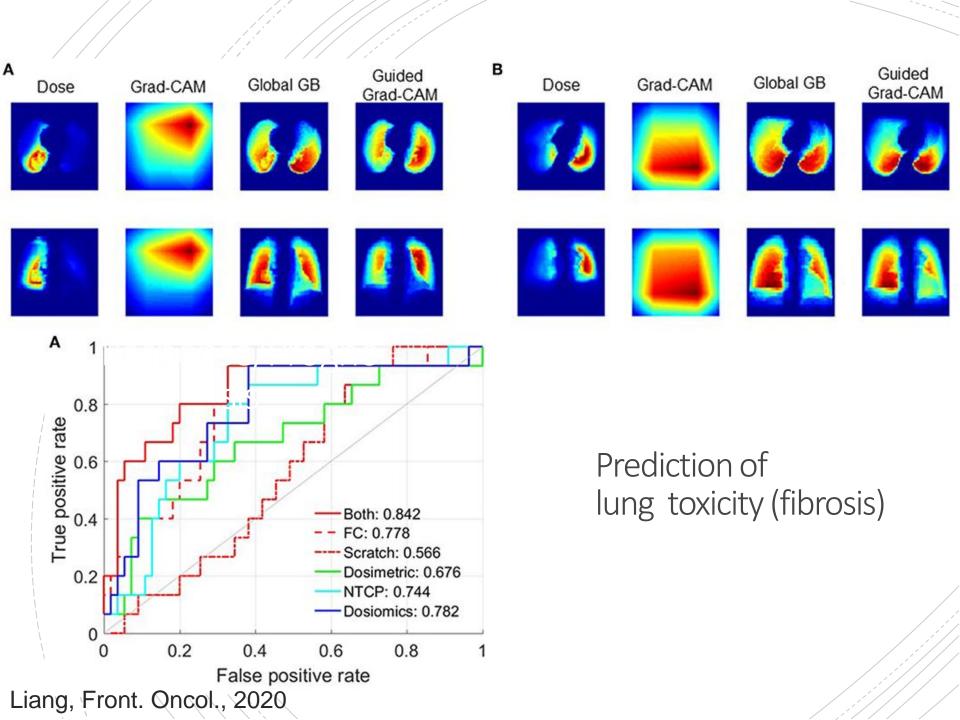
24.0 Gv

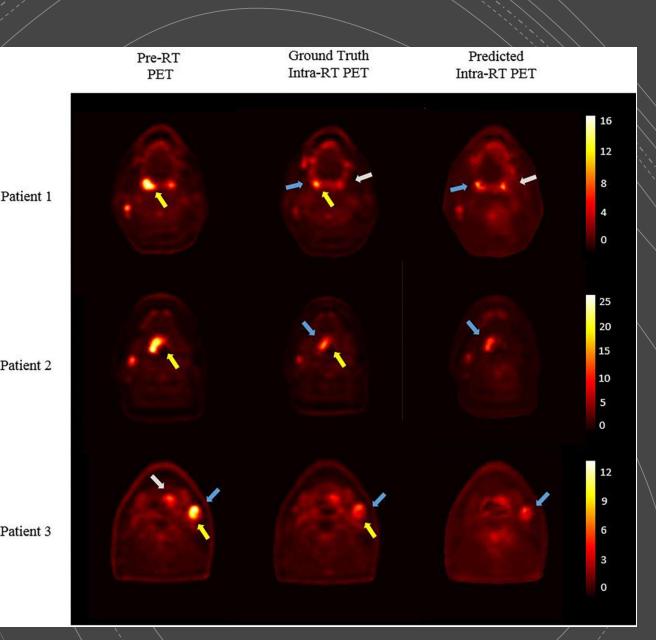
<u>Kunzel, Radiotherapy and Oncology, 2021</u>

### Prediction of acute toxicity to the rectum



Zhen, Phys. Med. Biol. 62 (2017) 8246

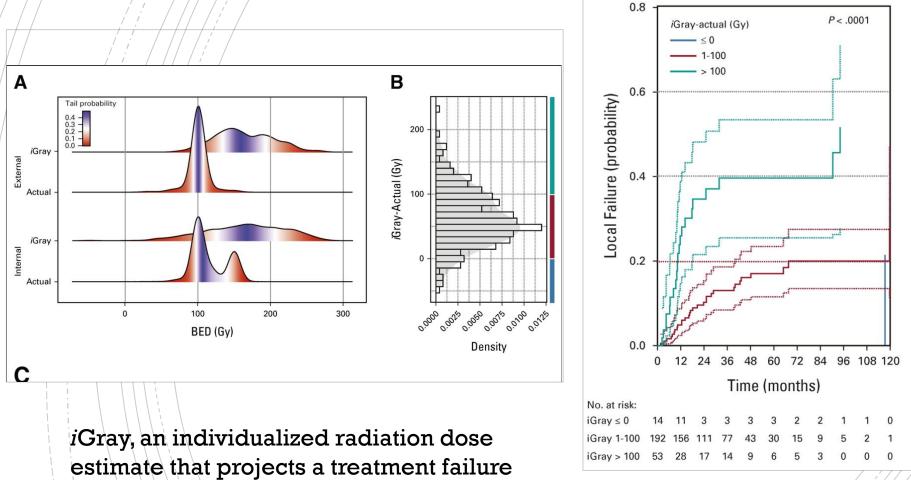




#### Prediction of tumor response in PET

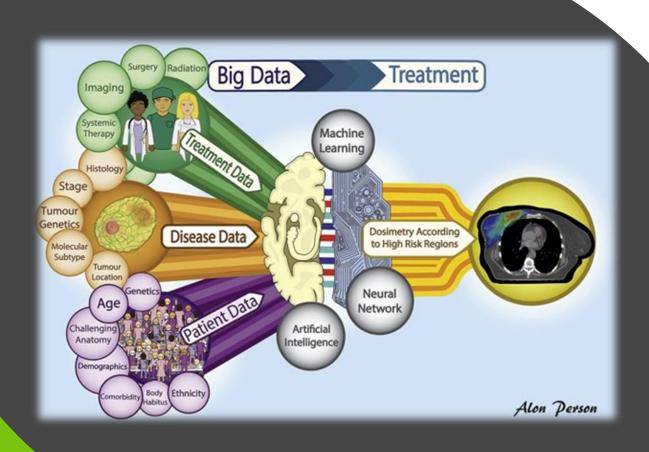
Wang, Front. Oncol., 2020

Dose prescription



probability of < 5% at 24 months

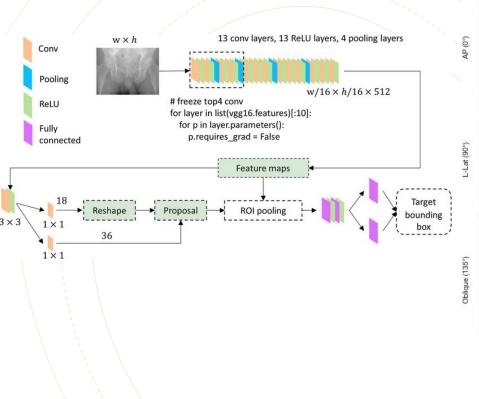
## Al guided treatment

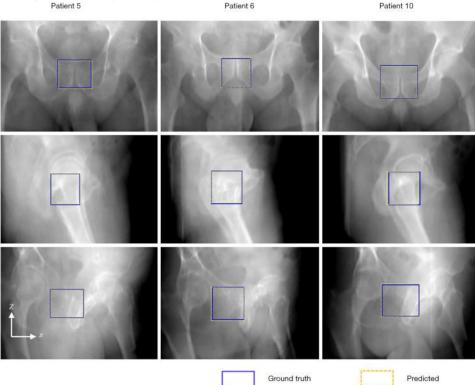


#### Radiotherapy treatment delivery



#### Target localization for IGRT



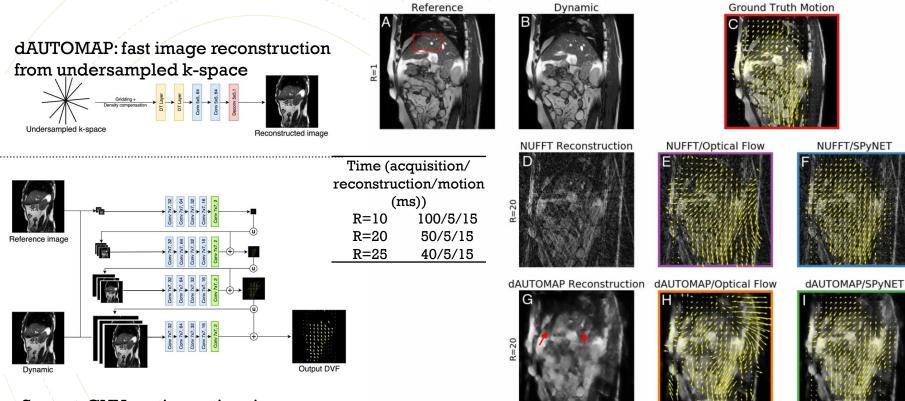


Patient 10

Zhao, Radiotherapy and Oncology, 2020

#### Deformable image registration

#### GAN

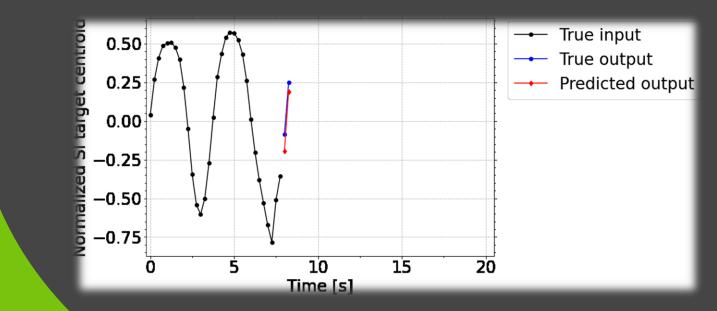


Spynet: CNN motion estimation

Terpstra et al, Physics in Medicine & Biology, Volume 65, Num

Comparison w traditional methods

Tumor centroid prediction: LSTM (Long Short Term Memory) networks (AI)



Tumor Tracking by Al

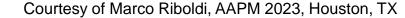


Gemelli

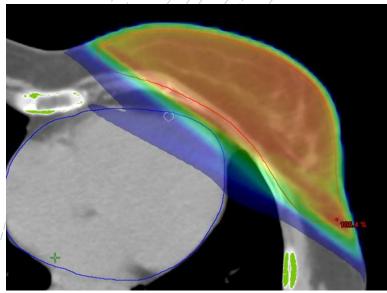
Università Cattolica del Sacro Cuore

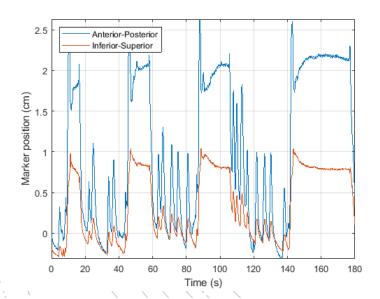
Fondazione Policlinico Universitario Agostino Gemelli IRCCS

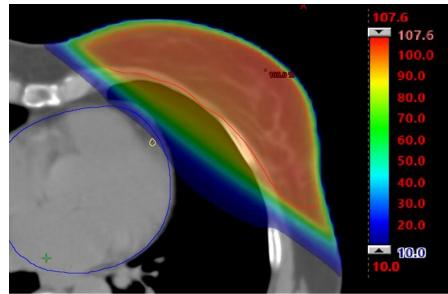
Advanced Radia Therapy

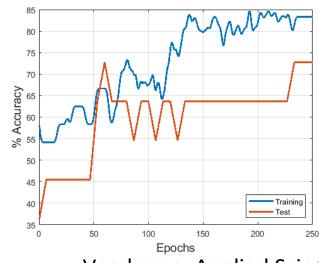


#### Patient selection for breath-hold RT



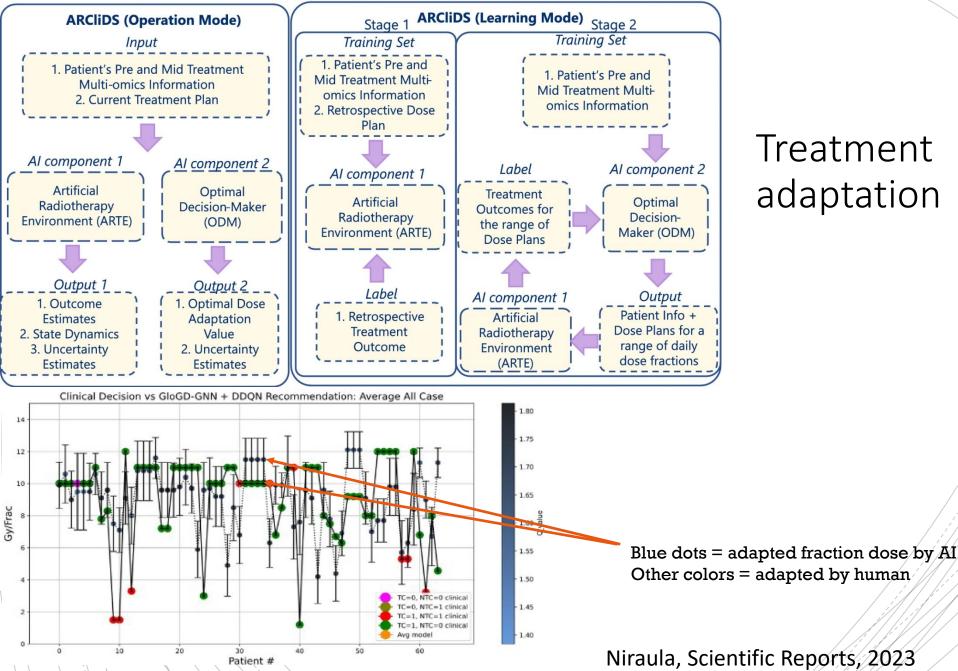






Vendrame, Applied Sciences, 2023

#### Adaptive Radiotherapy Clinical Decision Support system



#### Treatment adaptation

(b)

### Radiotherapy follow-up



## Recurrence +12m +3m +6m d lung injury +6m +12m +3m Benign changes

## Differentiation of relapse from fibrosis

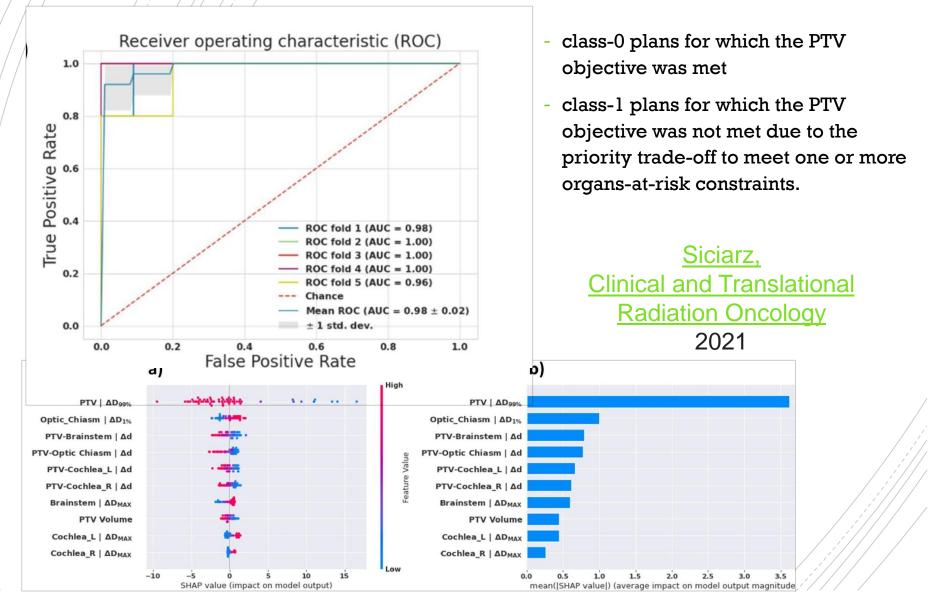
 Accuracy 76%–77% of radiomic features in crossvalidation

Mattonen et al. Med. Phys. 41 (3), March 2014

## Radiotherapy QA



#### Automated QA of treatment plans

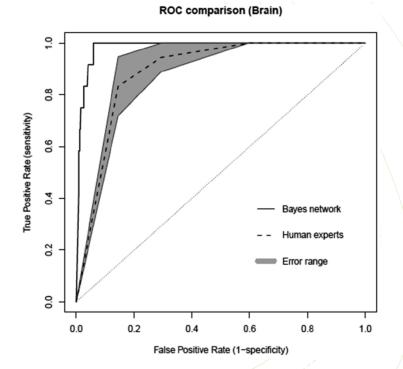


#### Quality check of treatment plans

Tx_Intent (site)	Technique	Modality	Dose_Ttl	Dose_Tx	numFields	Severity
Palliative (Brain)	Conformal Plan	×06	5400	200	6	7
Curative (Brain)	IMRT	×18	6000	200	12	2
Palliative (Breast)	PA	×18	2268	81	2	8

Table 1.	Example	potential	error	cases
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Note: Introduced errors are indicated in bold.

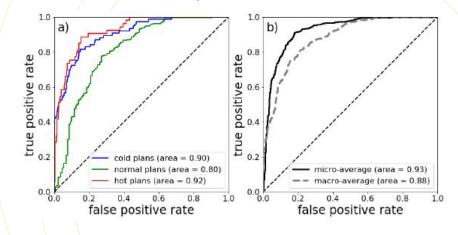


Alan M Kalet 2015 Phys. Med. Biol.

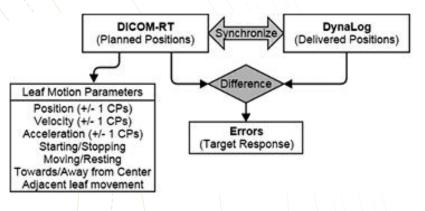
#### Automated QA of LINACs by AI

Prediction of results of Delivery Quality Assurance (DQA) of VMAT

Prediction of Delivery Quality Assurance of VMAT success/failure by ML



prediction of multi-leaf collimator positional errors Using dynalog files



A Granville et al 2019 Phys. Med. Biol. 64 095017

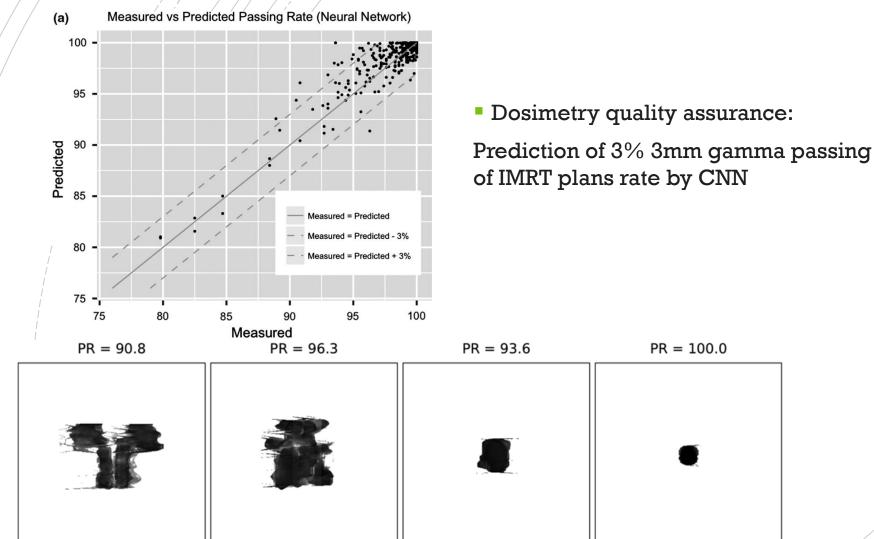
Carlson et al., Phys. Med. Biol. 61:2514.

References	QA Source	Data Source	ML Model	Task
Carlson et al. (2016)	DICOM_RT, Dynalog files	74 VMAT plans	Regression, Random Forest, Cubist	MLC Position Errors Detection
Li and Chan (2017)	Daily QA Device	5-year Daily QA Data	ANN Time-Series, ARIMA Models	Symmetry Prediction
Sun et al. (2018)	Ion Chamber	1,754 Proton Fields	Random Forrest, XGBoost, Cubist	Output for Compact Proton Machine
El Naqa et al. (2019)	EPID	119 Images from 8 Linacs	Support Vector Data Description, Clustering	Gantry Sag, Radiation Field Shift, MLC Offset
Grewal et al. (2020)	Ion Chamber	4,231 Proton Fields	Gaussian Processes, Shallow NN	Output and Patient QA Proton Machine
Osman et al. (2020)	log files	400 machine delivery log files	ANN	MLC Discrepancies during Delivery & Feedback
Chuang et al. (in press)	Trajectory log files	116 IMRT plans, 125 VMAT plans	Boosted Tree Outperformed LR	MLC Discrepancies during Delivery & Feedback
Zhao et al. (in press)	Water Tank Measurement	43 Truebeam PDD, Profiles	Multivariate Regression (Ridge)	Modeling of Beam Data

#### QA of radiotherapy

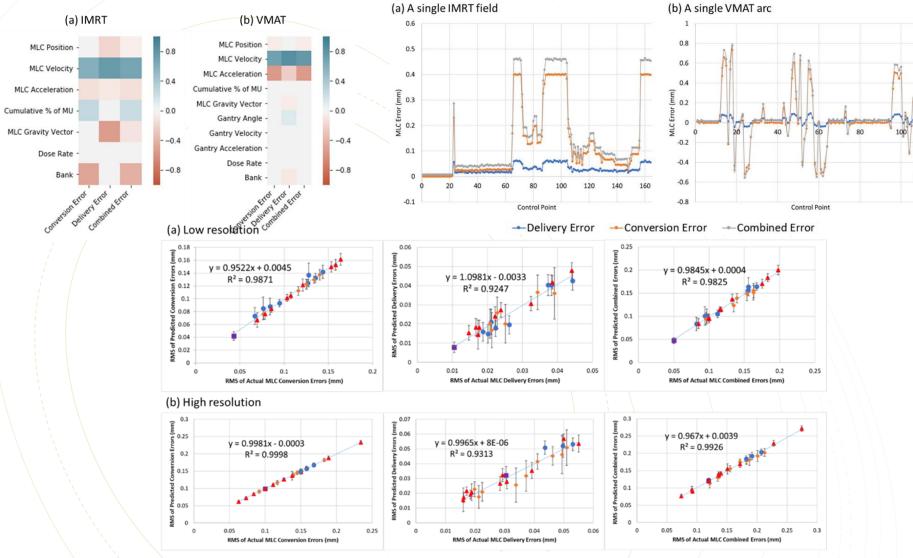
https://www.frontiersin.org/articles/10.3389/frai.2020.577620/full

## Prediction of gamma passing of IMRT plans



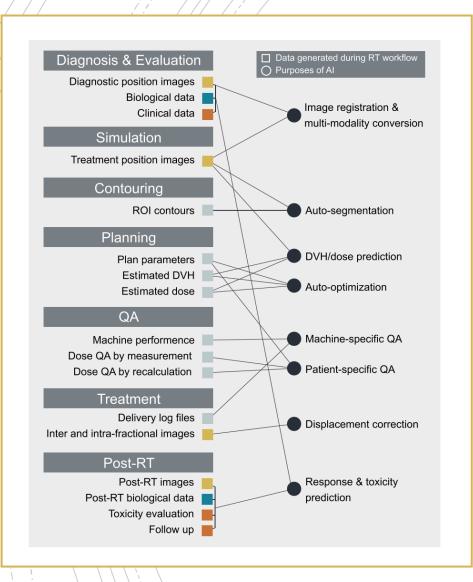
Interian Medical Physics, 2018

#### Prediction of errors in treatment delivery



Lay, SoftwareX, 2022

STX 
TB1 
TB2 
TB3



#### Summary

 Li, Seminars in Cancer Biology 2022

#### Selected bibliography

Introduction to AI:

 Shen, C.; Nguyen, D.; Zhou, Z.; Jiang, S.B.; Dong, B.; Jia, X. An Introduction to Deep Learning in Medical Physics: Advantages, Potential, and Challenges. *Phys.Med.Biol.* 2020, 65, 05TR01-6560/ab6f51, doi:10.1088/1361-6560/ab6f51.

Role of the Medical Physicist:

 Avanzo, M.; Trianni, A.; Botta, F.; Talamonti, C.; Stasi, M.; Iori, M. Artificial Intelligence and the Medical Physicist: Welcome to the Machine. *Applied Sciences* 2021, 11, 1691, doi:10.3390/app11041691.

Applications of AI to RT:

- Appelt, A.L.; Elhaminia, B.; Gooya, A.; Gilbert, A.; Nix, M. Deep Learning for Radiotherapy Outcome Prediction Using Dose Data - A Review. *Clin Oncol (R Coll Radiol)* 2022, 34, e87–e96, doi:10.1016/j.clon.2021.12.002.
- Vandewinckele, L.; Claessens, M.; Dinkla, A.; Brouwer, C.; Crijns, W.; Verellen, D.; van Elmpt, W. Overview of Artificial Intelligence-Based Applications in Radiotherapy: Recommendations for Implementation and Quality Assurance. *Radiother Oncol* 2020, *153*, 55–66, doi:10.1016/j.radonc.2020.09.008.