# Strengths, weaknesses and opportunities of artificial intelligence (AI) in radiation oncology

### Valerio Nardone

### Department of Precision Medicine, University of Campania «L. Vanvitelli», Naples



#### Disclosures

Per quanto concerne tale presentazione, dichiaro di non avere avuto alcuna relazione rilevante (diretta od indiretta) di tipo finanziario con alcuna compagnia farmaceutica negli ultimi 24 mesi che possa essere considerato un **conflitto di interesse.** 



Update degli Studi Practice Changing 2022

#### Outline

#### > Definition;

- Application in RadOnc;
- Highlights from ESTRO;
- Highlights from ASTRO;
- Other application;
- Summary and conclusions;



Update degli Studi Practice Changing 2022

In computer science, artificial intelligence (AI), is intelligence demonstrated by machines, in contrast to the *natural intelligence* displayed by humans and animals. The term AI is used to describe machines that mimic *cognitive* functions associated with human minds, such as *learning* and problem solving.



Update degli Studi Practice Changing 2022

#### **Other terms**

- Machine learning: it is an application of AI that provides systems the ability to aumatically learn and improve from experience without explicit programmation;
- Neural Networks: are computing systems inspired bu the biological neural networks and nodes called artificial neurons.
- Data mining: is the practice of esaming large databases to generate new informations;



**Artificial Intelligence in Factories** 



Japan ranked fourth in the world: In 2016, 303 robots were installed per 10,000 employees in the manufacturing industry.

#### Update degli Studi Practice Changing 2022

#### AI in Health

2019 & onwards

·Discovery and

development of

·Preclinical research

·Personalized Health

·And many more

### **Timeline of AI in health**

#### 1955

Term coined by John McCarthy. Founded as an academic discipline in 1956 in US.

#### Growth of microcomputer and new levels of network connectivity. AI systems in healthcare was designed to accommodate the absence of perfect data and build on the expertise of physicians.

1980s-1990s

#### 1960-1970

Produced first problemsolving program, or expert system, known as Dendral assisting to identifying bacteria and recommending antibiotics

#### 2010-2019

 Genomic sequencing databases

drugs

Care

- AI in electronic health record systems
- Natural language processing and computer vision,
- · Robot-assisted surgery. etc



Update degli Studi Practice Changing 2022

#### Digital Life Con la sua intelligenza artificiale ChatGPT scrive articoli scientifici a prova di... revisione umana

Oltre a produrre conversazioni complesse, ChatGPT può scrivere abstract di articoli scientifici e pezzi divulgativi che sembrano scritti da umani.

Special Reports > Exclusives

AI Passes U.S. Medical Licensing Exam

- Two papers show that large language models, including ChatGPT, can pass the USMLE

by Michael DePeau-Wilson, Enterprise & Investigative Writer, MedPage Today January 19, 2023

# Generative storytelling https://beta.tome.app/ has arrived.

Unlock your best work with Tome's AI-powered storytelling format.



**GPT** 

Update degli Studi Practice Changing 2022

#### Outline

- Definition;
- > Application in RadOnc;
- Highlights from ESTRO;
- Highlights from ASTRO;
- Other application;
- Summary and conclusions;



#### Update degli Studi Practice Changing 2022

2

### AI in Radiation Oncology



Huynh et al. Nat Rev Clin Oncol 2020

Update degli Studi Practice Changing 2022



Hype cycle. This figure features a hype cycle curve for three major innovations in radiation oncology (triangle: Monte Carlo; square: Inverse optimization/IMRT; circle: deep learning-based contouring). The curve depicts expectations by the target audience (those in radiation oncology and medical physics) as a function of time. Yellow, magenta, cyan, green, and blue portions of the curve denote "innovation trigger," "peak of inflated expectations," "trough of disillusionment," "slope of enlightenment," and "productivity plateau" regions, respectively.

The Emergence of Artificial Intelligence within Radiation Oncology Treatment Planning, Tetherton et al. Oncology 2021

#### Update degli Studi Practice Changing 2022

#### Use of AI in 2020



C.L. Brouwer et al. Physics and Imaging in Radiation Oncology 2020



Supplemental Table 1. Number of Radiation Oncology Departments using or preparing to use machine learning applications in clinical practice per country

Radiation Oncology Departments and				
Machine Learning Applications				
	Clinical	Preparing		
France	8			
Italy	7	5		
Netherlands	8	3		
Spain	7	2		
Australia	5	4		
Belgium	5	3		
United Kingdom	4	3		
Denmark	3	2		
Switzerland	3	3		
United States of Amerika	3	1		
Germany	2	2		
Norway	2	3		

Update degli Studi Practice Changing 2022

#### Outline

- Definition;
- Application in RadOnc;
- > Highlights from ESTRO;
- Highlights from ASTRO;
- Other application;
- Summary and conclusions;



Update degli Studi Practice Changing 2022

### AI: friend or foe

#### AI is a friend

- Time reduction for the radiotherapist (contours, planning),
- Improve accuracy and precision in RT treatments,
- Predict toxicity,
- Cost reduction.

#### Al is a foe

- " I may take your job ",
- Incorrect treatment decisions with incomplete or biased data,
- Manipulation of AI algorithms or steal patient data by hackers.





Update degli Studi Practice Changing 2022

1 1 1 1

### Al in Brachytherapy



ESTRO 2022	iysician point of view		Auditorium 11
LUCA TAGLIAFERRI - MD, PhD Artificial intelligen Current reality and	ce and brachythera perspectives	Post Post Post Post Post Post Post Post	ESTRO 2022 Virality
Gemelli 🚳 ART	Interventional Redatherapy INTERACTS Active Teaching School		Al a gatate
ESTRO 2022 The ph	ysicist point of view		Auditorium 11
Nicole Nesvacil Dept. for Radiation Oncology Medical University of Vienna Artificial Intelligenc reality A physic	ce and brachythera and perspectives cist's point of view	py: Current	ESTRO 2022 Rock NESVACI. Guistina
			Rika guesten
<b>EST<u>R0</u>20</b>	22 6 <mark>-10 May</mark>	<b>2022</b> en, Denmark	
	and the second second		

Update degli Studi Practice Changing 2022

### AI in the RT workflow



#### Update degli Studi Practice Changing 2022





Update degli Studi Practice Changing 2022

#### Outline

- Definition;
- Application in RadOnc;
- Highlights from ESTRO;
- Highlights from ASTRO;
- Other application;
- Summary and conclusions;



Update degli Studi Practice Changing 2022



Emotional intelligence is defined as **the ability to understand and manage your own emotions, as well as recognize and influence the emotions of those around you**. The term was first coined in 1990 by researchers John Mayer and Peter Salovey, but was later popularized by psychologist Daniel Goleman.

Update degli Studi Practice Changing 2022

### Impact of AI on Quality of Care, Clinical Practice and Training

- Variation in quality exists, but regionalization is NOT the solution;
- Al can reduce unrewarding tasks that consume time/cognitive burden;
- AI can also inform or facilitate strategies to enhance quality (simulation training for complex cases, patient selection for escalation of clinical care);
- There will be pluses and minuses, but change is certain;
- Success will come through networking within us and across disciplines;



Erin Gillespie



Update degli Studi Practice Changing 2022

### Deep learning in digital histopathology for prostate cancer

- Al derived prognostic biomarkers provide personalized risk estimates, that when grouped allows more streamlined communication;
- ArteraAI MMAI prognostic tool identifies 6-fold more low-risk patients than NCCN (safe omitting of ADT with RT, with a NNT>25);
- Prognostic biomarkers help with shared-decision making to avoid futile treatment intensification;
- Use of AI tools leveraging digital pathology improves prognostication, enabling us to determine the optimal treatment plan for the single patient (precision medicine);



Jonathan Tward



Update degli Studi Practice Changing 2022

#### Current Progress of Machine Learning in Radiation Oncology

Please tell me how concerned you are about the use of Al in medicine for each of the following:

- 1. That my health information will be kept confidential
- 2. That AI will make the wrong diagnosis
- That AI will mean I spend less time with my doctor
- 4. That Al will increase my healthcare costs





Sanjay Aneja

For each of the following, please tell me how comfortable you would feel with AI doing some of the things your doctor usually does:

- 1. Reading your chest x-ray
- 2. Diagnosing pneumonia
- 3. Telling you that you have pneumonia
- 4. Recommending your antibiotic
- 5. Diagnosing cancer
- 6. Telling you that you have cancer





Update degli Studi Practice Changing 2022

Exploring ethical challenges in RadOnc AI

- While waivers can be ethical and pragmantic solutions, patients have no idea that AI is being used in research or care involving them;
- If for minimal risk for quality systems we can rely on good ML practice, for higher risk we necessitate prospective informed consent!
- Umbrella consent needed to inform patients;
- Additional disclosure duties: patient access to information about specific AI algorithms used in their care;



Subha Perni



Update degli Studi Practice Changing 2022

#### Outline

- Definition;
- Application in RadOnc;
- Highlights from ESTRO;
- Highlights from ASTRO;
- > Other application;
- Summary and conclusions;



#### Cost-effectiveness analysis in oncology with AI

JAMA Oncology | Brief Report

#### Long-term Effect of Machine Learning-Triggered Behavioral Nudges on Serious Illness Conversations and End-of-Life Outcomes Among Patients With Cancer A Randomized Clinical Trial JAMA 2023

Christopher R. Manz, MD; Yichen Zhang, PhD; Kan Chen, MA; Qi Long, PhD; Dylan S. Small, PhD; Chalanda N. Evans, BS; Corey Chivers, PhD; Susan H. Regli, PhD; C. William Hanson, MD; Justin E. Bekelman, MD; Jennifer Braun, MHA, RN, BSN; Charles A. L. Rareshide, MS; Nina O'Connor, MD; Pallavi Kumar, MD, MPH; Lynn M. Schuchter, MD; Lawrence N. Shulman, MD; Mitesh S. Patel, MD, MBA; Ravi B, Parikh, MD, MPP

#### healthcare

Article Economics of Artificial Intelligence in Healthcare: Diagnosis vs. Treatment

Narendra N. Khanna<sup>1</sup>, Mahesh A. Maindarkar<sup>2,3</sup>, Vijay Viswanathan<sup>4</sup>, Jose Fernandes E Fernandes<sup>5</sup>,

#### Healthcare 2023

MDPI

**CONCLUSIONS AND RELEVANCE** In this randomized clinical trial, a machine learning-based behavioral intervention and behavioral nudges to clinicans led to an increase in SICs and reduction in end-of-life systemic therapy but no changes in other end-of-life outcomes among outpatients with cancer. These results suggest that machine learning and behavioral nudges can lead to long-lasting improvements in cancer care delivery.

Update degli Studi Practice Changing 2022

#### Genomic analysis in oncology with Al

Validation of a 40-gene expression profile test to predict metastatic risk in localized high-risk cutaneous squamous JAAD 2021 cell carcinoma

Ashley Wysong, MD, MS,<sup>a</sup> Jason G. Newman, MD,<sup>b</sup> Kyle R. Covington, PhD,<sup>c</sup> Sarah J. Kurley, PhD,<sup>c</sup> Sherrif F. Ibrahim, MD, PhD,<sup>d</sup> Aaron S. Farberg, MD,<sup>e,f</sup> Anna Bar, MD,<sup>g</sup> Nathan J. Cleaver, DO,<sup>h</sup> Ally-Khan Somani, MD, PhD,<sup>i</sup> David Panther, MD,<sup>j</sup> David G. Brodland, MD,<sup>j</sup> John Zitelli, MD,<sup>j</sup> Jennifer Toyohara, MD,<sup>k</sup> Ian A. Maher, MD,<sup>1</sup> Yang Xia, MD,<sup>m</sup> Kristin Bibee, MD,<sup>n</sup> Robert Griego, MD,<sup>o</sup> Darrell S. Rigel, MD,<sup>p</sup> Kristen Meldi Plasseraud, PhD,<sup>c</sup> Sarah Estrada, MD,<sup>q,r</sup> Lauren Meldi Sholl, MS,<sup>q</sup> Clare Johnson, RN,<sup>q</sup> Robert W. Cook, PhD,<sup>c</sup> Chrysalyne D. Schmults, MD, MSCE,<sup>s</sup> and Sarah T. Arron, MD, PhD<sup>t</sup>

#### Seminars in Cancer Biology 88 (2023) 187–200

AL.

Contents lists available at ScienceDirect
Seminars in Cancer Biology

journal homepage: www.elsevier.com/locate/semcancer

Artificial intelligence-based multi-omics analysis fuels cancer precision medicine

Xiujing He<sup>1</sup>, Xiaowei Liu<sup>1</sup>, Fengli Zuo, Hubing Shi, Jing Jing



- JAAD 2021 Development and validation of a nomogram incorporating gene expression profiling and clinical factors for accurate prediction of metastasis in patients with cutaneous melanoma following Mohs micrographic surgery

Ryan B. Thorpe, MD,<sup>a</sup> Kyle R. Covington, PhD,<sup>b</sup> Hillary G. Caruso, PhD,<sup>b</sup> Ann P. Quick, PhD,<sup>b</sup>



Update degli Studi Practice Changing 2022

### Genomic analysis in oncology with AI: the risk for RADONC

#### Development and Validation of a Genomic Profile for the Omission of Local Adjuvant Radiation in Breast Cancer JCO 2023

Martin Sjöström, MD, PhD<sup>1,2</sup>; Anthony Fyles, MD<sup>3</sup>; Fei-Fei Liu, MD<sup>3</sup>; David McCready, MD<sup>3</sup>; Wei Shi, MSc<sup>3</sup>; Katrina Rey-McIntyre, MBA, BSc<sup>3</sup>; S. Laura Chang, PhD<sup>4</sup>; Felix Y. Feng, MD<sup>2</sup>; Corey W. Speers, MD, PhD<sup>5</sup>; Lori J. Pierce, MD<sup>5</sup>; Erik Holmberg, PhD<sup>6</sup>; Mårten Fernö, PhD<sup>1</sup>; Per Malmström, MD, PhD<sup>1,7</sup>; and Per Karlsson, MD, PhD<sup>6</sup>

### **BUT CONSIDER THIS**

### Testicular Cancer in the Cisplatin Era: Causes of Death and Mortality Rates in a Population-Based Cohort

Ragnhild Hellesnes, MD<sup>1,2</sup>; Tor Åge Myklebust, PhD<sup>3,4</sup>; Sophie D. Fosså, MD, PhD<sup>4,5,6</sup>; Roy M. Bremnes, MD, PhD<sup>1,2</sup>; Ása Karlsdottir, MD, PhD<sup>7</sup>; Øivind Kvammen, MD, PhD<sup>8</sup>; Torgrim Tandstad, MD, PhD<sup>9,10</sup>; Tom Wilsgaard, PhD<sup>11</sup>; Helene F. S. Negaard, MD, PhD<sup>5</sup>; and Hege S. Haugnes, MD, PhD<sup>1,2</sup>

radiotherapy in patients age ≥ 70 with T1N0 ER/PR + HER2- breast cancer treated with breast conserving surgery and endocrine therapy Neil Chevli •Wagar Hague • Kevin T. Tran • ... Sandra S. Hatch • E. Brian Butler • Bin S. Teh A 🗠 •

Show all authors

21-Gene recurrence score predictive for prognostic benefit of

Published: June 27, 2022 • DOI: https://doi.org/10.1016/j.radonc.2022.06.013 • 🜔 Check for updates



...

#### Genomic analysis in oncology with AI: the opportunity for RADONC

)rk 1

se is

...

Drew Moghanaki @DrewMoghanaki · 16h In risposta a @theabzlab Congratulations on pushing us into the future.

I noticed a very high event rate in your cohort, suggesting these data will need to be validated in a healthier cohort to be applicable to operable pts.



ts in the other lung or pleura ory were excluded from univar ultivariate analyses because o w sample size and variability in ogic subtypes. Local failure was d as radiographic progression 1 cm of the planning target e to maintain a consistent :ion of local/marginal failure in I trials of SBRT.<sup>1,13,14</sup> Prescripti

仚

111 153

#### Drew Moghanaki @DrewMoghanaki · 16h In risposta a @DrewMoghanaki e @theabzlab

This is a MUST READ paper for anyone coding local failures after SBRT. It shows how easy, but not how often, it is for coders to be fooled and overcall LF. doi.org/10.1016/j.ijro...



Update degli Studi Practice Changing 2022

#### Things to remember: AI performance depends on the TRAINING



Update degli Studi Practice Changing 2022

#### Genomic analysis in oncology with AI: the opportunity for RADONC

Journal Pre-proof

### Red Journal 2022

Genomic classifiers in personalized prostate cancer radiotherapy approaches – a systematic review and future perspectives based on international consensus

Simon K.B. Spohn MD, Cédric Draulans MD, Amar U. Kishan MD, Daniel Spratt MD, Ashley Ross MD, PhD, Tobias Maurer Prof. MD, Derya Tilki Prof., MD, Alejandro Berlin MD, Pierre Blanchard MD, PhD, Sean Collins MD, Peter Bronsert MD, Ronald Chen MD, PhD, Alan Dal Pra MD, Gert de Meerleer MD, Prof., Thomas Fade MD, Prof. Karin Haustermans MD, Prof.



Published in final edited form as: *J Thorac Oncol.* 2021 March ; 16(3): 428–438. doi:10.1016/j.jtho.2020.11.008.

#### Personalizing Radiotherapy Prescription Dose Using Genomic Markers of Radiosensitivity and Normal Tissue Toxicity in Non-Small Cell Lung Cancer

Jacob G. Scott<sup>1,2,†,\*</sup>, Geoff Sedor<sup>2,†</sup>, Jessica A. Scarborough<sup>1,2</sup>, Michael W. Kattan<sup>3</sup>, Jeffrey Peacock<sup>4</sup>, G. Daniel Grass<sup>4</sup>, Eric A. Mellon<sup>5</sup>, Ram Thapa<sup>6</sup>, Michael Schell<sup>6</sup>, Anthony Waller<sup>7</sup>, Sean Poppen<sup>8</sup>, George Andl<sup>7</sup>, Jamie Teer<sup>6</sup>, Steven A. Eschrich<sup>6</sup>, Thomas J. Dilling<sup>4</sup>, William S. Dalton<sup>9</sup>, Louis B. Harrison<sup>4</sup>, Tim Fox<sup>7</sup>, Javier F. Torres-Roca<sup>3,\*</sup>

# Pan-cancer prediction of radiotherapy benefit using genomic-adjusted radiation dose (GARD): a cohort-based pooled analysis

Jacob G Scott, DPhil <sup>†</sup> • Geoffrey Sedor, MD <sup>†</sup> • Patrick Ellsworth, BA • Jessica A Scarborough, MS •

Kamran A Ahmed, MD • Daniel E Oliver, MD • et al. Show all authors • Show footnotes

Published: August 04, 2021 • DOI: https://doi.org/10.1016/S1470-2045(21)00347-8

Check for updates

Update degli Studi Practice Changing 2022

#### Outline

- Definition;
- Application in RadOnc;
- Highlights from ESTRO;
- Highlights from ASTRO;
- Other application;
- Summary and conclusions;



Update degli Studi Practice Changing 2022

### SUMMARY

- In the next years we will see many different application of AI in RadOnc workflow;
- Current applications focus on contouring, planning, adaptive;
- Future development in other areas (genomic profiling, radiomics) could have an high impact on RadOnc;
- Ethical challenges need to be solved;
- A big effort of RadOnc community is needed to train and validate AI approaches (networking is the key);



Update degli Studi Practice Changing 2022

#### **CONCLUSIONS**

# Don't be afraid of the future







3DRT

IMRT



VMAT











Update degli Studi Practice Changing 2022

# There is no SOUL in the MACHINE. Only in front of it.



# Without you, it's just DATA.

#### Update degli Studi Practice Changing 2022



Università degli Studi della Campania *Laigi Vanvitelli* 



