Interpretable Artificial Intelligence in Medicine: What Else?

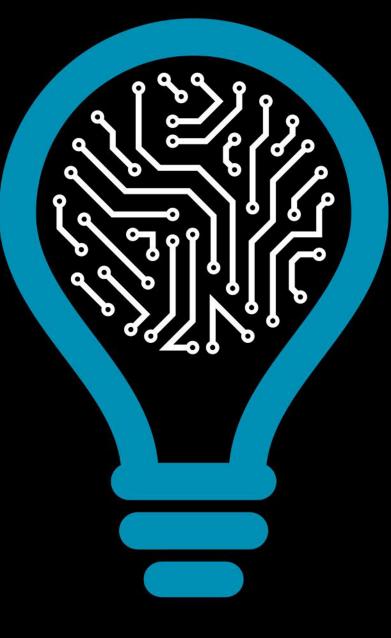
XIV Symposium on Bioengineering | Porto (Portugal) Made in Bio | March 18, 2023

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Outline

1. Interpretable Artificial Intelligence: The Bright Side of the Force

2. What's so interesting about medical data?

3. Towards the development of a Transparent New World

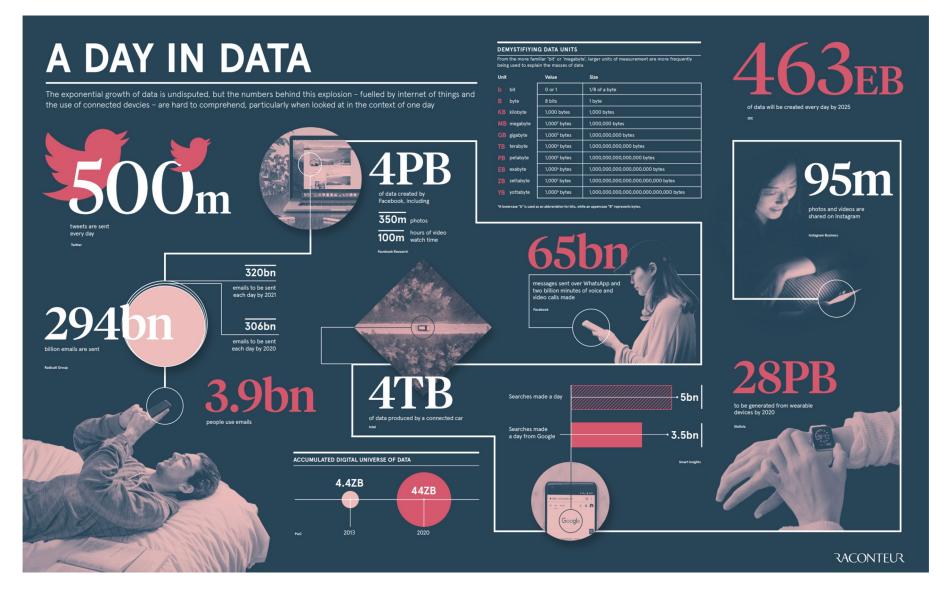


1. Interpretable Artificial Intelligence: The Bright Side of the Force

Nowadays, we are constantly generating data^[1]

- The paradigm is changing: most of the daily tasks and services can now be performed with the aid of digital applications or gadgets
- High-tech companies such as Google, Facebook, Netflix or Amazon have access to huge amounts of data from several data sources and users:
 - This phenomenon suggests that the *business of data* will become a significant sector of the global economy^[2]
 - There are several open-source data sets with millions of entries (e.g., ImageNet^[3])
- Data is referred as **the new oil**^[4]
 - The main impact on humanity is related **to the way data can improve our lives**
 - A proper management process of the "dark side" of data must be implemented, but the advances in data fuels are worth the effort

Take a look: A Day in the Wonderful World of Data^[1, 2, 3]



We have more computational power than ever

- The fundamental concepts of artificial intelligence and deep neural networks have been around since 1940^[1]
 - Frank Rosenblatt proposed one of the first approaches to the design and training of artificial neural networks: the Perceptron^[2]
- The development of powerful computer processing units (CPUs) and the leveraging of the graphical processing units (GPUs)^[3] for computation allowed the training of deep and complex algorithms in "human time"

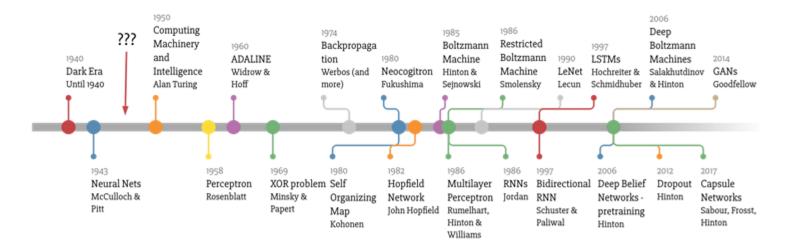


Figure - A (tentative) deep learning timeline (Image from [1])

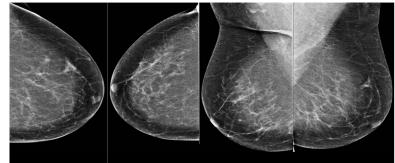
Sources: [1] https://towardsdatascience.com/a-weird-introduction-to-deep-learning-7828803693b0, [2] Frank Rosenblatt "The perceptron: A probabilistic model for information storage and organization in the brain.",
[3] https://www.nvidia.com/en-us/

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Technology has been *challenging* human performance...

- There are, at least, two popular events that created a revolution in the History of AI:
 - In 1997, IBM's Deep Blue beated the Chess World Champion Garry Kasparov^[1]
 - In 2016, Google's DeepMind AlphaGo learned to play Go alone (i.e., through reinforcement learning policies) and beat the Go World Champion Lee Sedol^[2]
- The two events above are examples of the (virtually) unlimited boundaries of the application of artificial intelligence to our daily lives
 - In 2020, Google's DeepMind published a paper in *Nature* suggesting that "its model was able to spot cancer in de-identified screening mammograms with fewer false positives and false negatives than experts"^[3, 4]

Figure - Medical Image Analysis: Mammograms (Image from [4])



Everything seems good except for the lack of transparency

- The increase of available computational power and the democratised access to a huge amount of data has leveraged the development of novel artificial intelligence (AI) algorithms and their applications
- Deep learning techniques have been challenging human performance at some specific tasks such as cancer detection in biomedical imaging^[1] or machine translation in natural language processing^[2]
- However, most of these models work as black boxes (i.e., their internal logic is hidden to the user) that receive data and output results without justifying their predictions in a human understandable way^[3]

No worries! We are working on that!

- Even if the models achieve high performances, it is not trivial to assure that they are learning features that are relevant for that domain (i.e., black box behavior)
 - Machine learning models are good at extracting correlations
- While this may not be an issue in several domains (e.g., recommendation systems), in others, it is of utmost importance that the system is capable of transparently showing the reasons behind its decisions (e.g., healthcare)

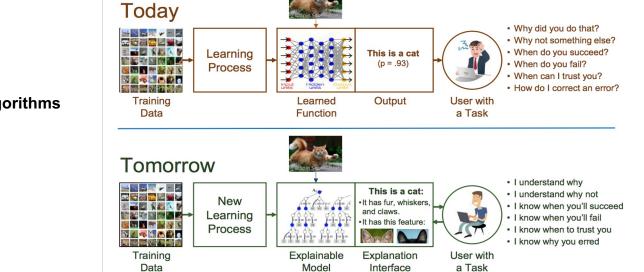


Figure - The future of machine learning algorithms (Image from [1])

Explain it like a Human: Interpretability is the key!

- Interpretability is a concept that results from the interaction between several definitions
 - The degree to which a human can understand the cause of a decision^[1]
 - The degree to which a human can **consistently predict the model's result**^[2]
- Interpretable machine learning is also related to the "extraction of relevant knowledge from a machine-learning model concerning relationships either contained in data or learned by the model"^[3]
- Intuitively, the higher the degree of interpretability of a model, the higher the likelihood of a user comprehending its predictions^[4]
- "Humans have a mental model of their environment that is updated when something unexpected happens. This update is performed by finding an explanation for the unexpected event"^[4]



2. What's so interesting about medical data?

Medical data is multimodal, and that is awesome

- In the clinical context, it is common to combine several image modalities during the decision making process (e.g. computed tomography, electroencephalography, magnetic resonance imaging, positron emission tomography)
- Recently, a comprehensive study^[1] on data fusion strategies for image classification and segmentation reported that the **network trained with multi-modal images showed superior performance** compared to networks trained with single-modal images, and that **performing image fusion within the network** (e.g., fusing at convolutional or fully connected layers) is generally better than fusing images at the network output (e.g., voting)^[1]

Multimodality means that data fusion will play a key role in our lives

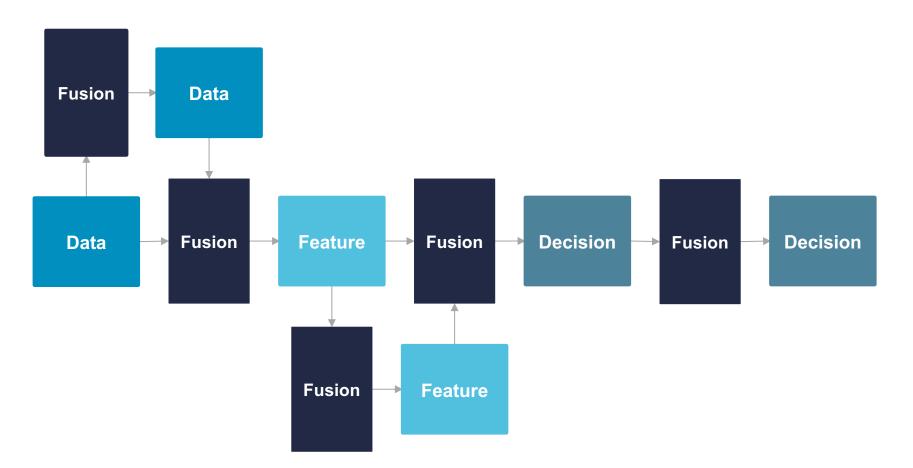


Figure: Different strategies of data fusion^[1]

The medical world needs human-understandable explanations

- The black box behavior of deep learning models does not help decision-makers to have a clear understanding of their inner-functioning, thus preventing them to diagnose errors and potential biases or deciding when and how much to rely on these models^[1]
- There has been a huge effort into the development of post-model strategies to explain the behavior of black box models, however, the outputs of these algorithms are prone to subjective evaluation, may be misleading^[2] or fooled^[1]
- Besides, we agree that just being able to obtain explanations is not enough and that we rather need to take into account at the development stage that these methods must respect specific constraints that give them the capability of generating humanunderstandable explanations and make decisions based on such premises^[3]

What do you see?^[1]

TABLE 12. Example of LRP and DeepLIFT *post-hoc* saliency maps for an image of the MIMIC-CXR data set with the label 0 correctly classified as 0 by all models.

	DENSENET			RESNET		
Original Image	No Attention	+ SE Layer	+ CBAM Layer	No Attention	+ SE Layer	+ CBAM Layer
DeiT (LRP)	No Attention	+ SE Layer	+ CBAM Layer	No Attention	+ SE Layer	+ CBAM Layer

The healthcare tech market is full of opportunities and requires interpretable artificial intelligence

- Interpretability is already playing its role in the pipelines of machine learning deployment: researchers and developers are using interpretability techniques to validate and debug their models before deployment^[1, 2]
- Regarding the availability of end-user software that contain machine learning algorithms for medical applications, we point to the popularity of:
 - Software for the analysis of volumetric medical images
 - Software for the development and creation of **DICOM pipelines and servers**
 - Software for the **annotation** and **segmentation** of medical images
 - Software for the **automatic classification** of medical images
 - Software for the automatic analysis of electronic health records of patients to generate diagnosis and recommendations



3. Towards the development of a Transparent New World

Responsible AI relies on fundamental principles

- **Responsible AI** is a framework that guides how we should address the challenges around artificial intelligence from both an **ethical, technical and legal** point of view^[1]
 - We must resolve ambiguity for where responsibility lies if something goes wrong!
- This framework relies on fundamental principles^[2]:
 - Accountability
 - Interpretability
 - Fairness
 - Safety
 - Privacy

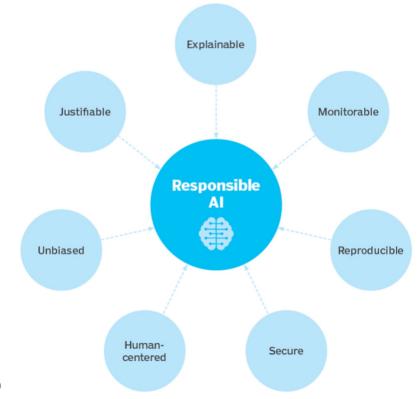


Figure - Responsible AI (Image from [1])

We need to design ethical and fair algorithms

- To facilitate trust (and increase transparency) in AI algorithms it is important to **ensure a priori that these models are interpretable**, and understand how decisions are made in the clinical context
- On the other hand, it is important to understand what the algorithms are already learning and to **evaluate the quality of such explanations** (e.g., understand if the algorithms are extracting relevant features for the clinical context)
- A different dimension of the application of AI in sensitive domains such as healthcare is the development of ethical and fair algorithms^[1]
 - This strategy is supported by the new European Union's General Data Protection Regulation (EU-GDPR)^[2] which advises that these algorithms should be able to explain their decisions for the sake of transparency

While keeping an attentive eye on the technologies that are shaping our lives

- Many entities are already leveraging their data sources to optimize their inner processes or to develop new services or products, thus enabling them to achieve a substantial competitive advantage^[1]
- In the healthcare context, systems and algorithms need to go through a continuous pipeline of validation and error assessment
- Hence, it is **reasonable to accept that these technologies may need to be calibrated** to the data sources of the institutions that are integrating them into their information systems and that these algorithms **may have a continuous learning policy over time**
- Moreover, to assure transparency, accountability and accessibility, new regulatory frameworks will have to be developed to allow model adaptations that enable optimal performance while ensuring reliability and patient safety^[2]

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