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# Coalitions intégratives pour une utilisation de confiance de l'intelligence artificielle en imagerie médicale

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MIAM: Midi de l'Intelligence Artificielle pour la Médecine

Benoit Macq – [benoit.macq@uclouvain.be](mailto:benoit.macq@uclouvain.be)

[www.pilab.be](http://www.pilab.be)

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# Artificial Intelligence

## ARTIFICIAL INTELLIGENCE

A program that can sense, reason, act, and adapt

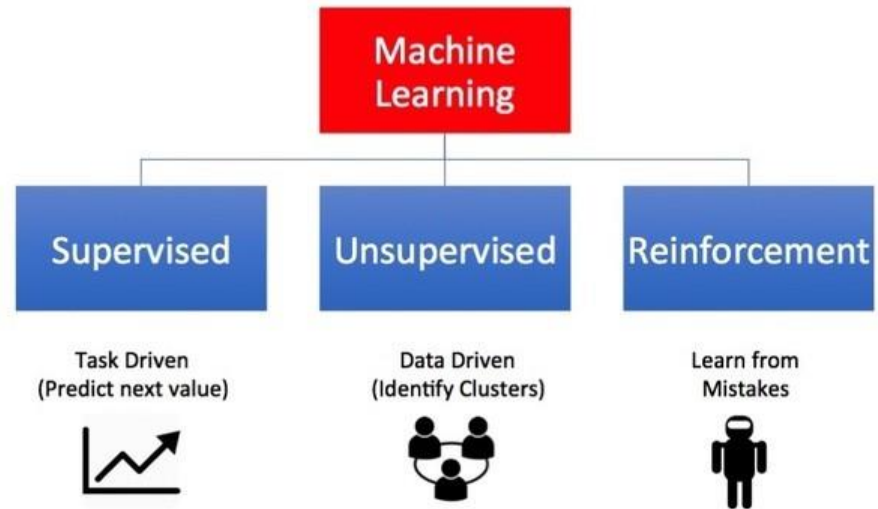
## MACHINE LEARNING

Algorithms whose performance improve as they are exposed to more data over time

## DEEP LEARNING

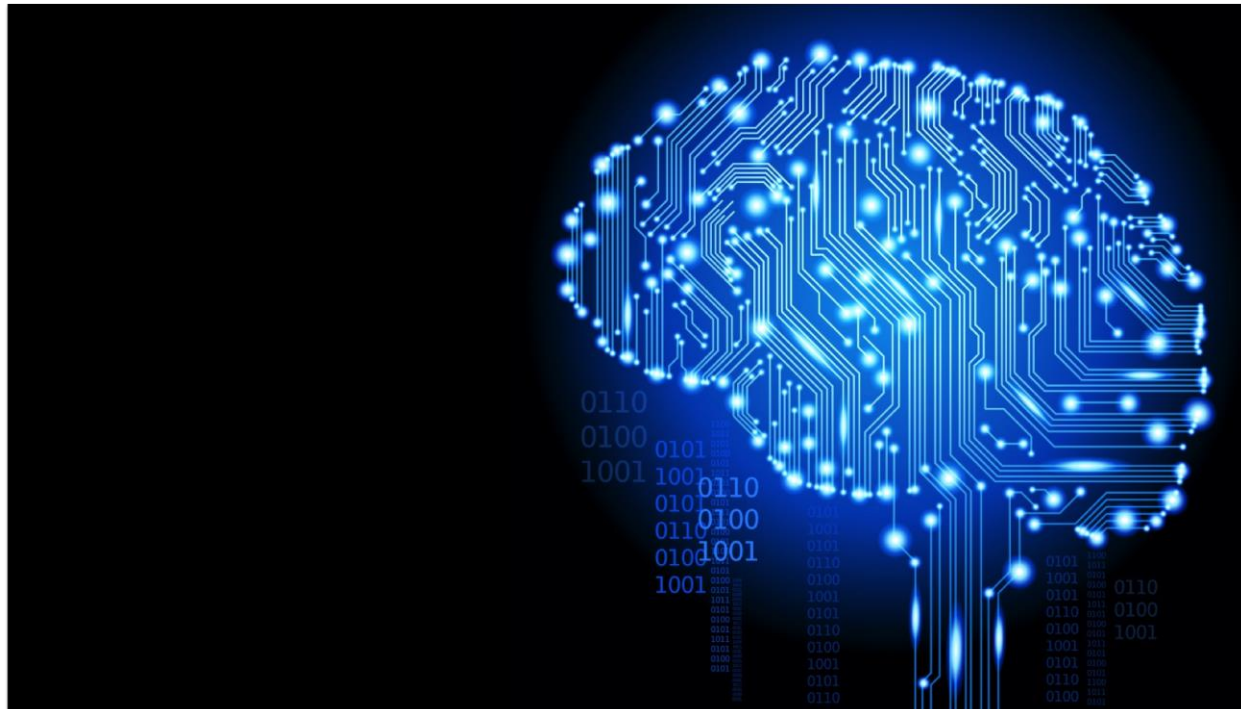
Subset of machine learning in which multilayered neural networks learn from vast amounts of data

## Types of Machine Learning



# Deep learning (e.g. MILA-Montréal)

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# Deep learning success stories

## Gaming



AlphaGo beat (4-1) world champion Lee Sedol (*March 2016*)

## Self-driving car



Vehicles have driven 1.6m km  
Fautive in 1 crash (June 2015)

## Image recognition



ImageNet Challenge: classify 1.2m high-res. images  
U. of Toronto team reaches 17% top-5 error rate (2012)

## Why not medical imaging ?



MIT technology 50 smartest companies  
6000 lung cancer diagnoses  
50% more accurate than human radiologists



Watson Health medical imaging collaborative  
15 health systems, medical centers and imaging comp.  
Data from ~300m patients



Applying machine learning to RT planning for H&N cancer  
Objective: segmentation process 4 hours → 1 hour

ImagX, BidMed, ... Telemis, Intuitim, DNAlytics, Oncoradiomics

# Image-based decision in the previous millenium

IEEE TRANSACTIONS ON BIOMEDICAL ENGINEERING, VOL. 43, NO. 10, OCTOBER 1996

## Morphological Feature Extraction for the Classification of Digital Images of Cancerous Tissues

Jean-Philippe Thiran,\* *Student Member, IEEE*, and Benoît Macq, *Member, IEEE*

**Abstract**— This paper presents a new method for automatic recognition of cancerous tissues from an image of a microscopic section. Based on the shape and the size analysis of the observed cells, this method provides the physician with nonsubjective numerical values for four criteria of malignancy. This automatic approach is based on mathematical morphology, and more specifically on the use of Geodesy. This technique is used first to remove the background noise from the image and then to operate a segmentation of the nuclei of the cells and an analysis of their shape, their size and their texture. From the values of the extracted criteria, an automatic classification of the image (cancerous or not) is finally operated.

### I. INTRODUCTION

OBJECTIVE analysis of microscopic images of cells and

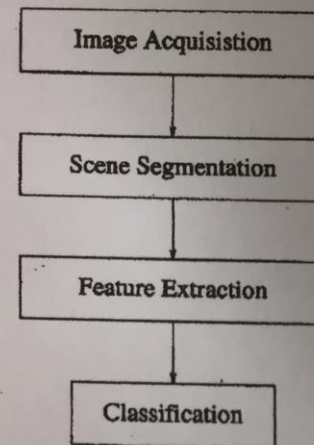
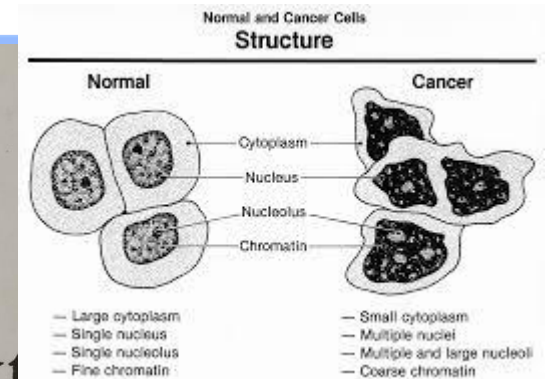
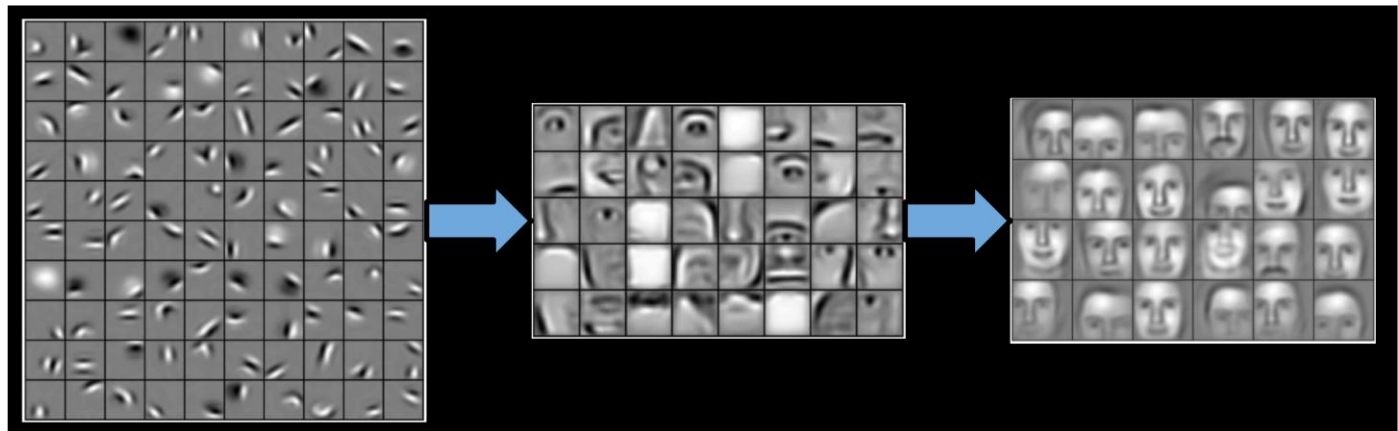
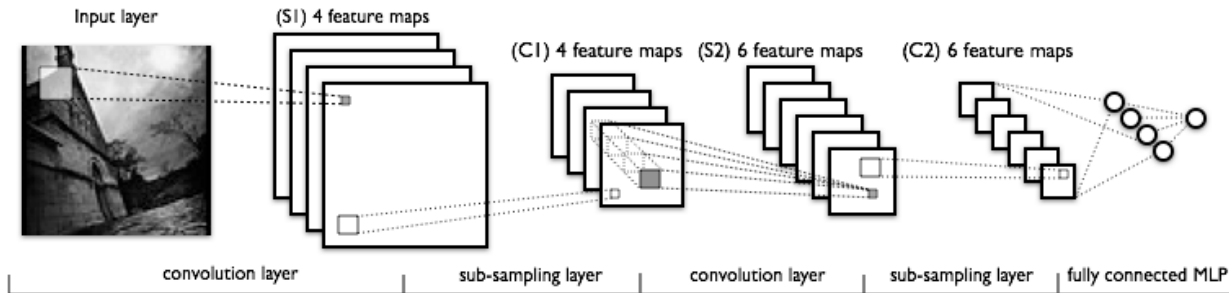


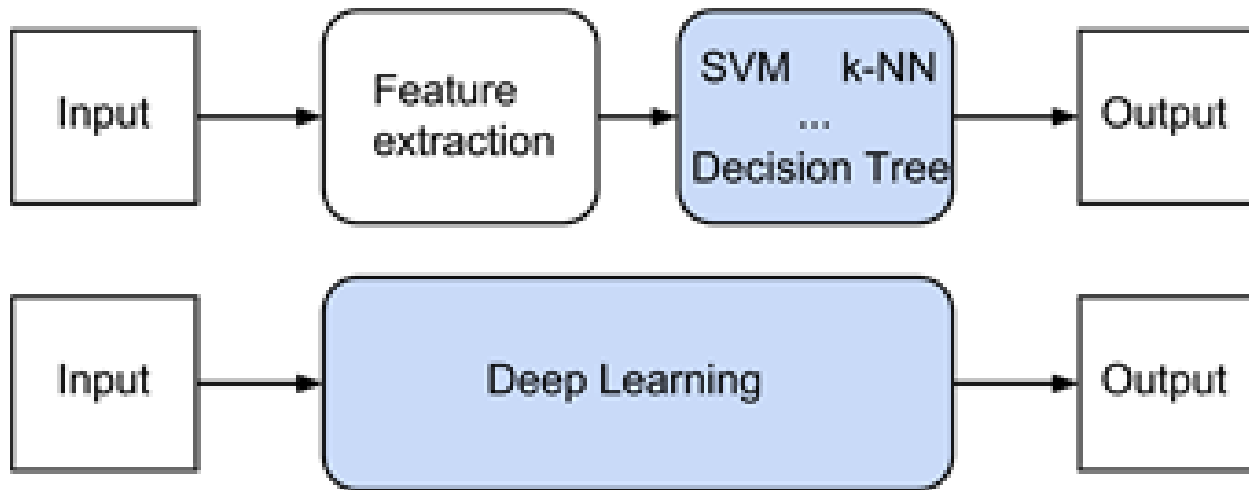
Fig. 1. General structure of the method.

# Deep Learning high predictive power (e.g. facial recognition)



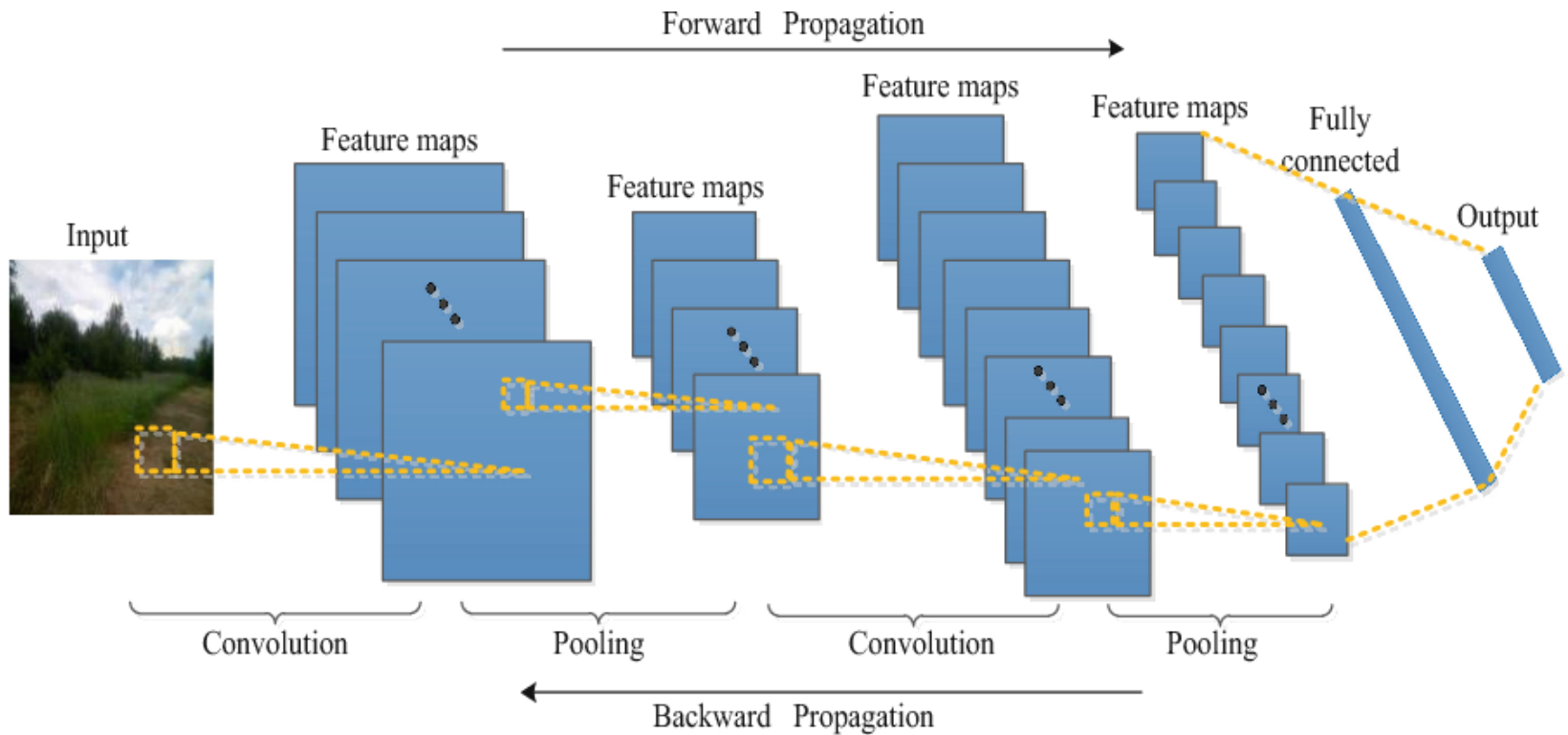
# The move based on the high predictivity power of deep learning

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# Convolutional Neural Networks (CNN)



**Fig. 1.** The CNN structure



# Pro and cons of Deep Learning

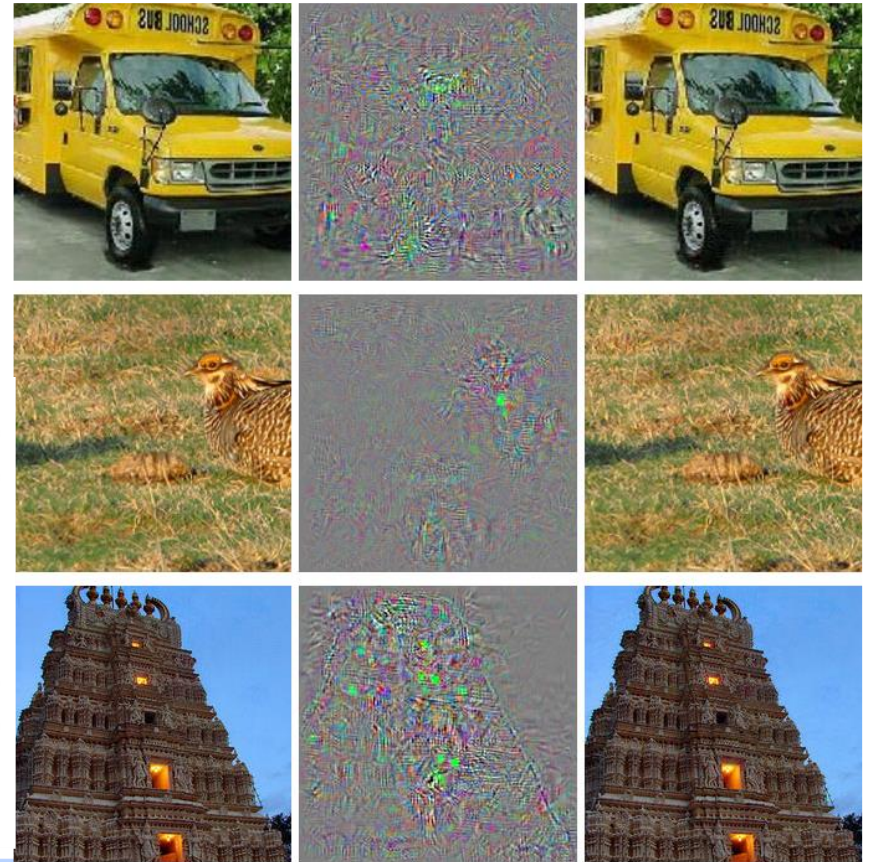
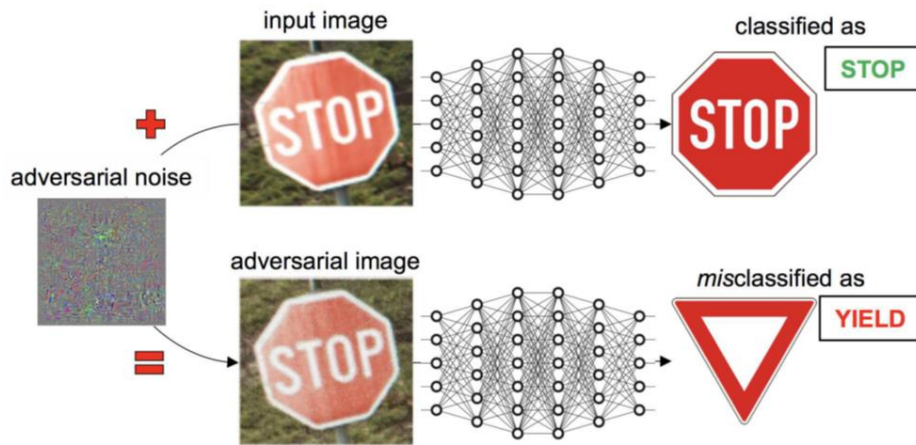
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- Unique structure (CNN) for many problems
  - « Generalisable » with regards the training set
  - There exist a lot of opensource tools
  - But: need for large annotated training sets
  - But: lack of explainability of the deep features and their co-action
  - But: lack of actionability
-

# And ....

- Intriguing properties of neural networks  
(C. Szegedy et al. )

<https://www.pluribus-one.it/research/sec-ml/wild-patterns>

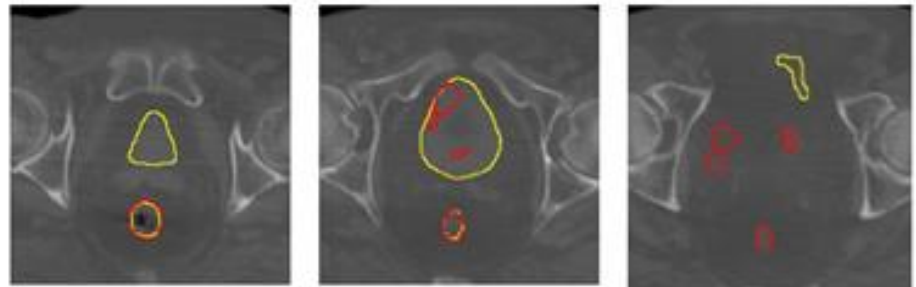


# Deep Learning has an outstanding accuracy in difficult problems but hard to explain outliers

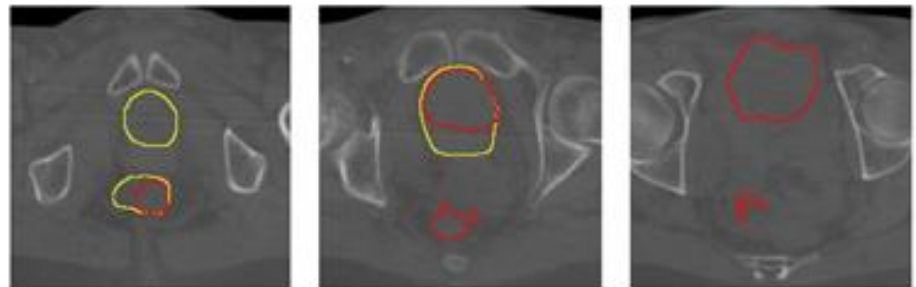
## U-net fails for two patients

For bladder

Patient 30  
(DSC = 0.124)



Patient 44  
(DSC = 0.211)



# L'Intelligence Artificielle et les radiologues

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**« Les radiologues qui utiliseront l'IA remplaceront ceux qui ne l'utilisent pas »**

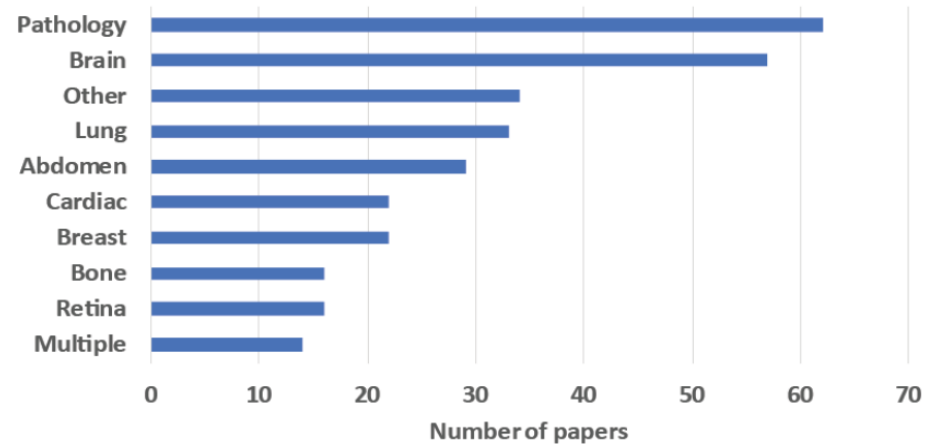
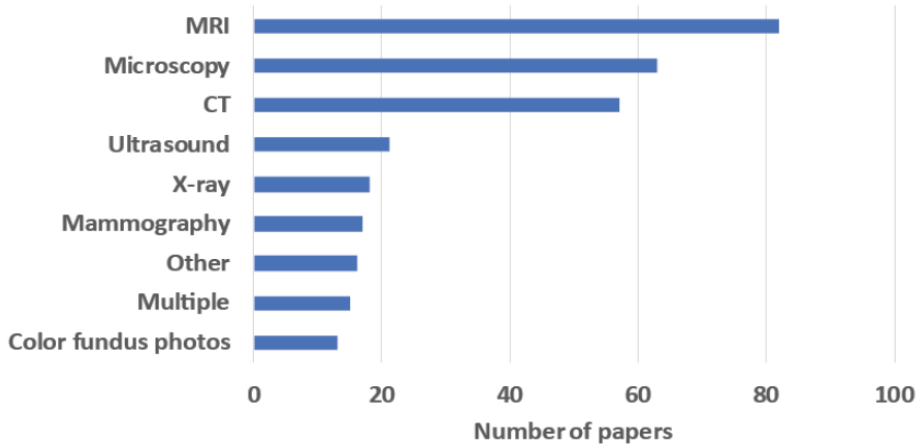
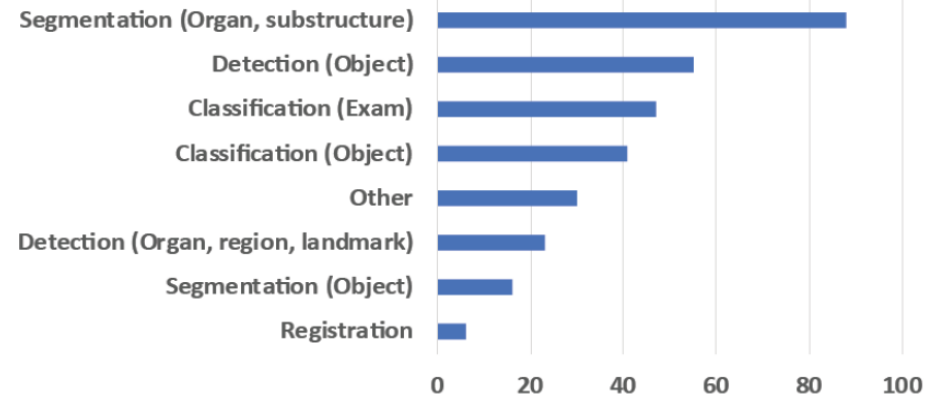
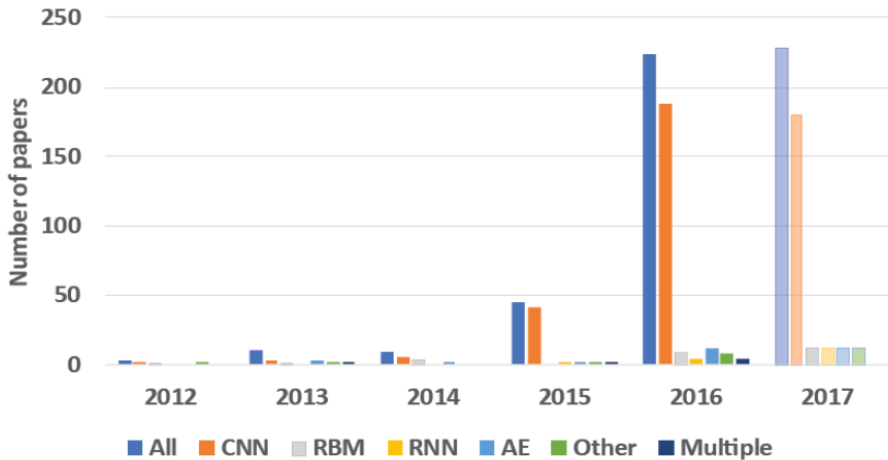
1. « Unpredictable » outliers (reliability ?)
2. Explainability of the decision ?
3. Actionability and commitment
4. Data privacy (blockchained distributed learning)
5. Evolution of expertise



**« Prédire n'est pas comprendre »**

Explainability of algorithms (GDPR)

# Deep learning in medical imaging



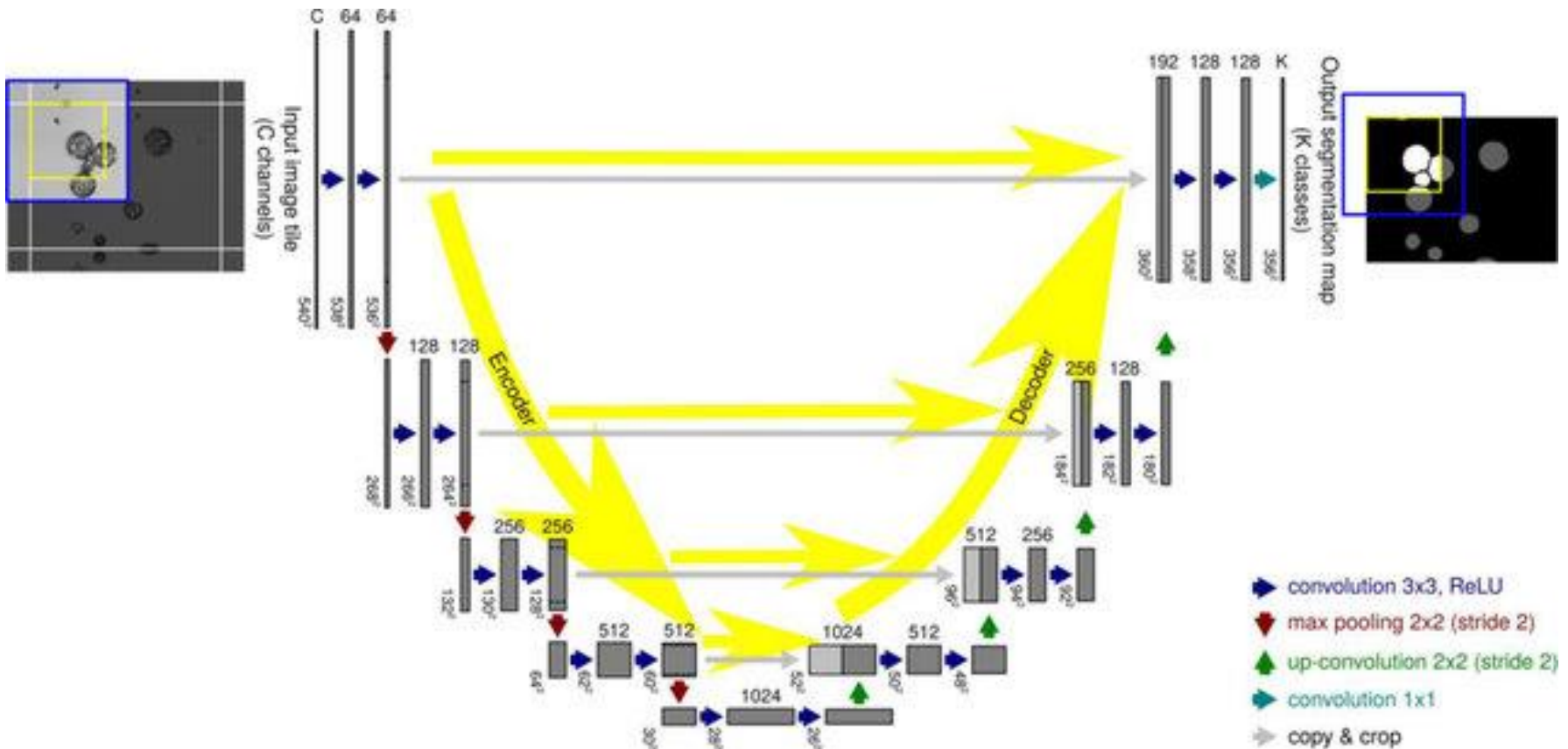
Litjens, *et al.* a survey on deep learning in medical image analysis. *Med Image Anal.* 2017

# Advanced deep learning

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- U-Net segmentation
- Generative Adversarial Networks (GANs)
- Deep Reinforcement learning

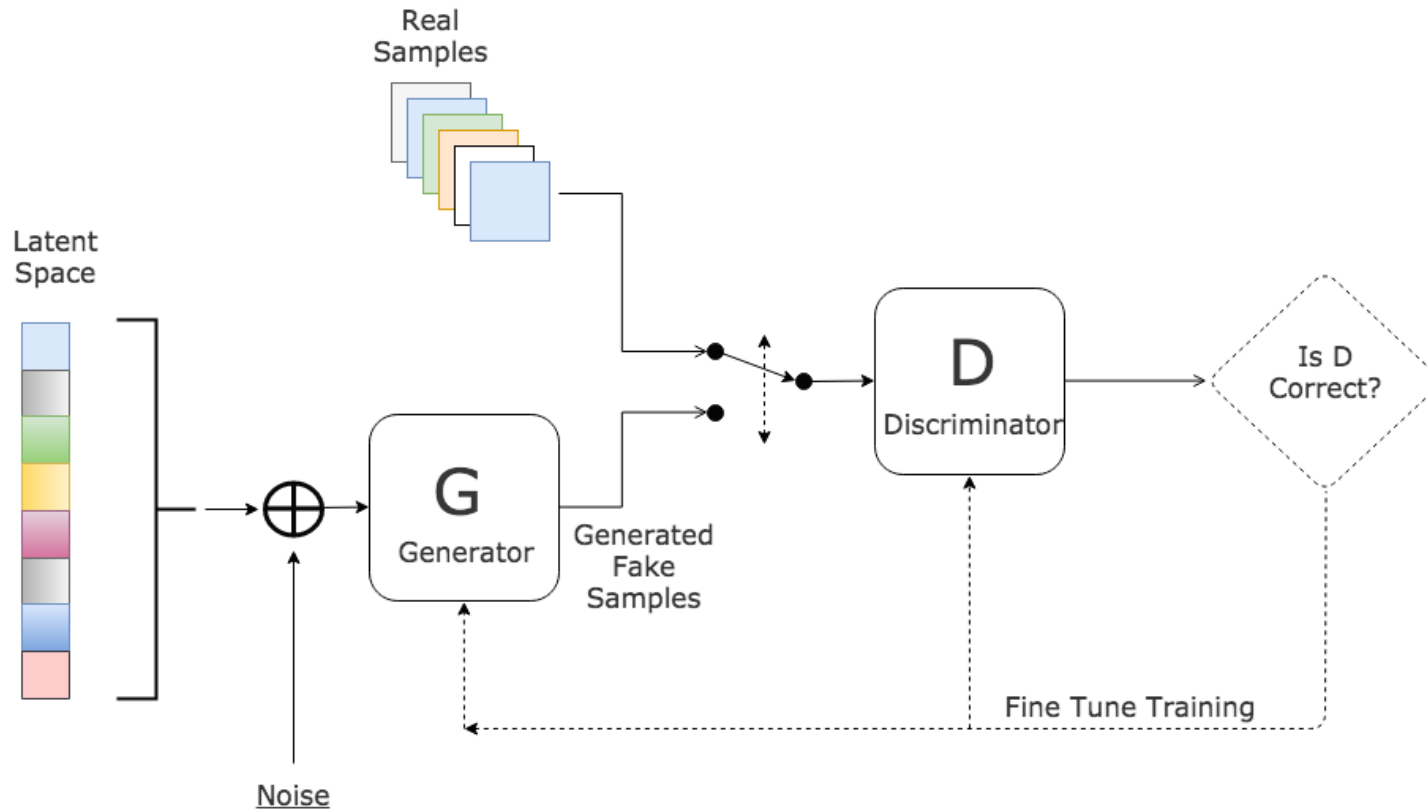
# U-Net Segmentation





# GANs

## Generative Adversarial Network



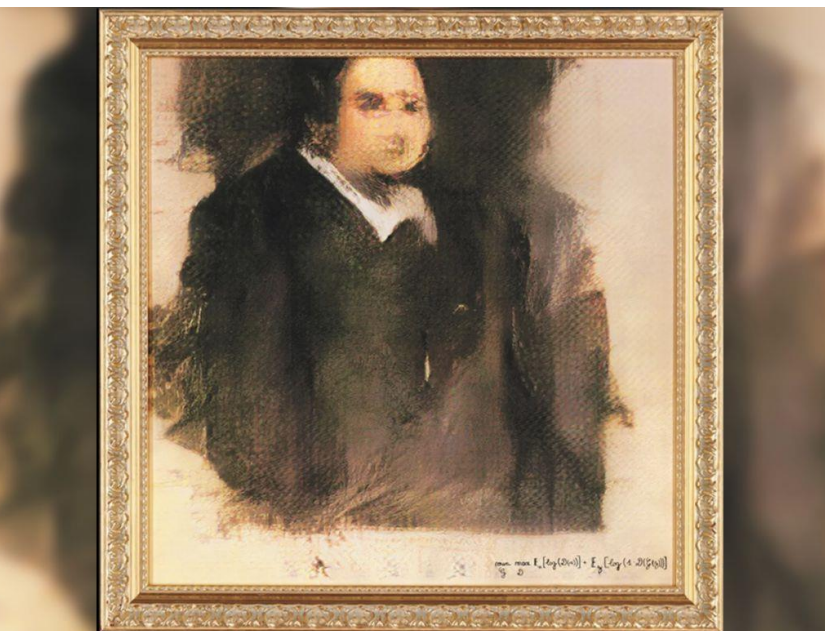
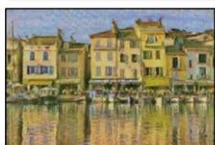
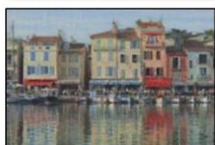
Input

Monet

Van Gogh

Cezanne

Ukiyo-e



source

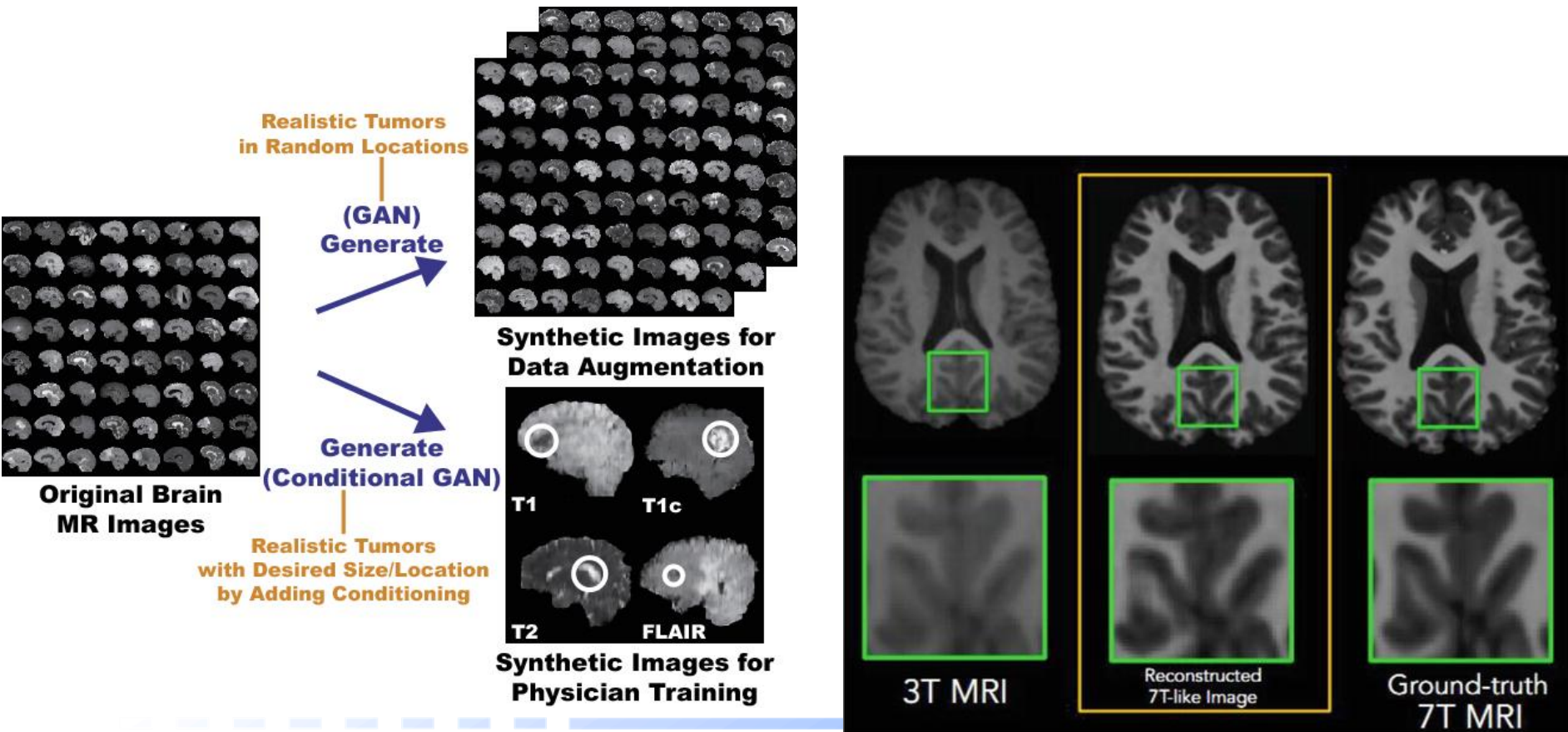


destination





# Use of GAN in radiology



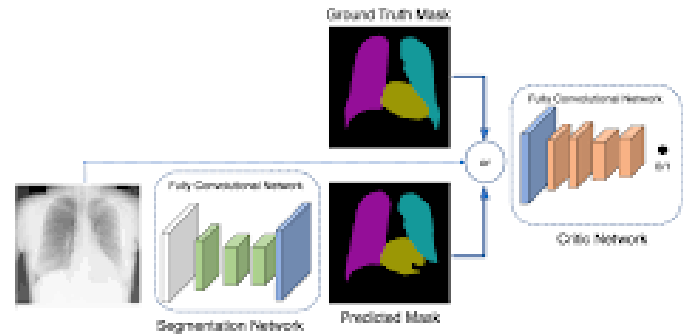
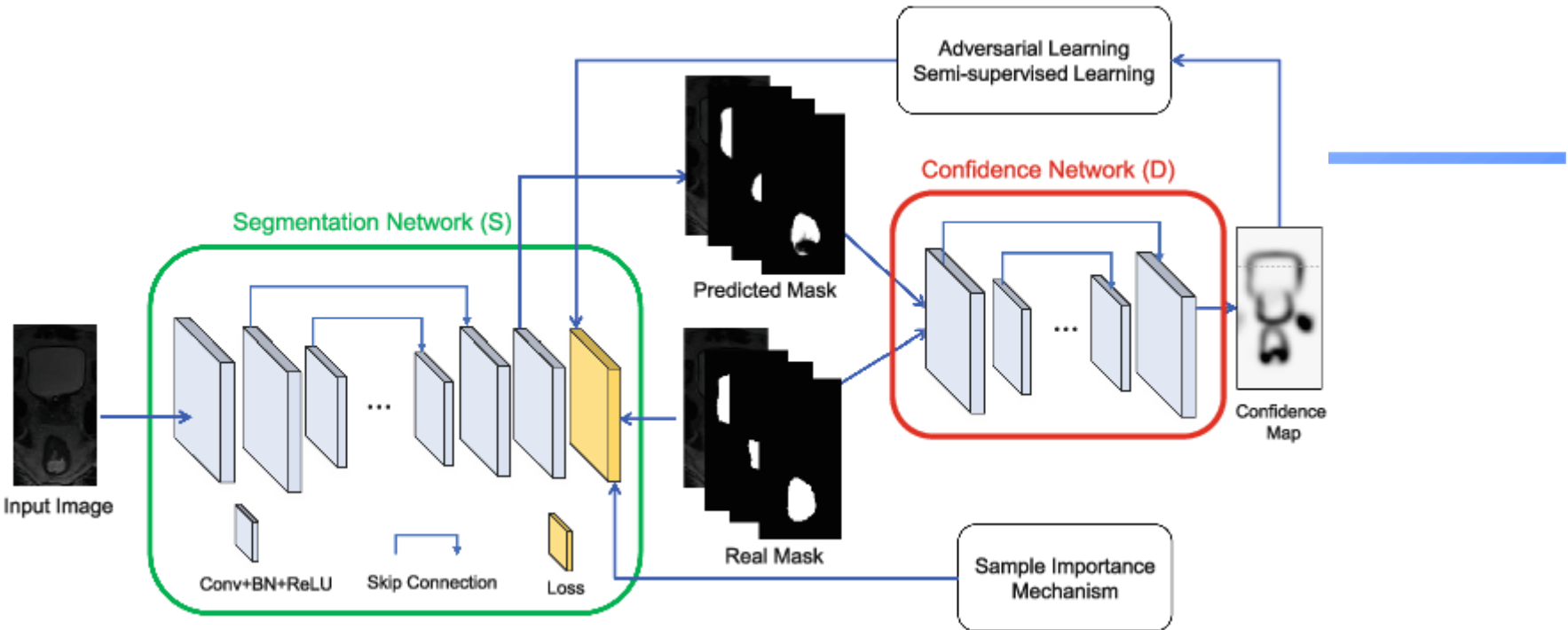
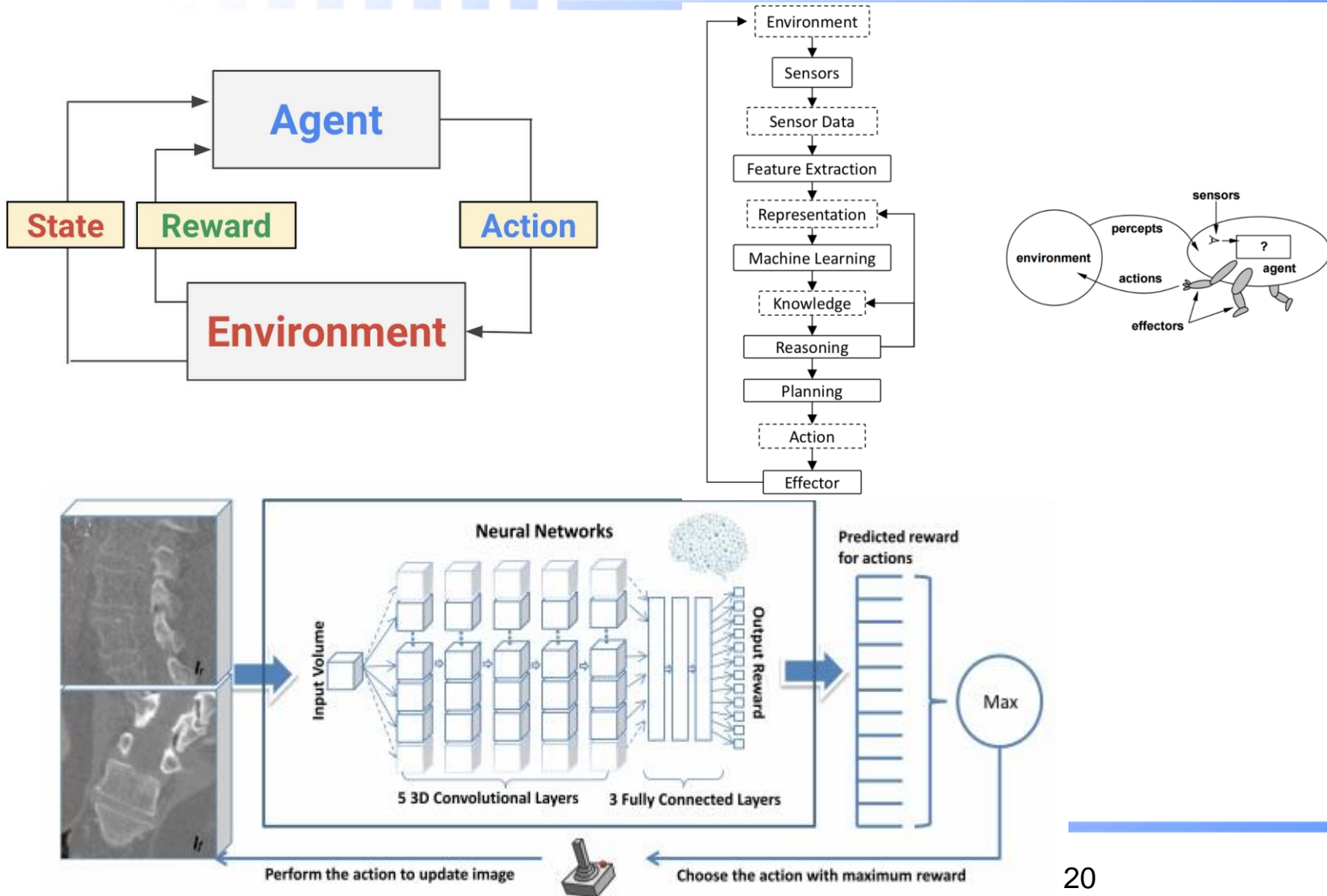
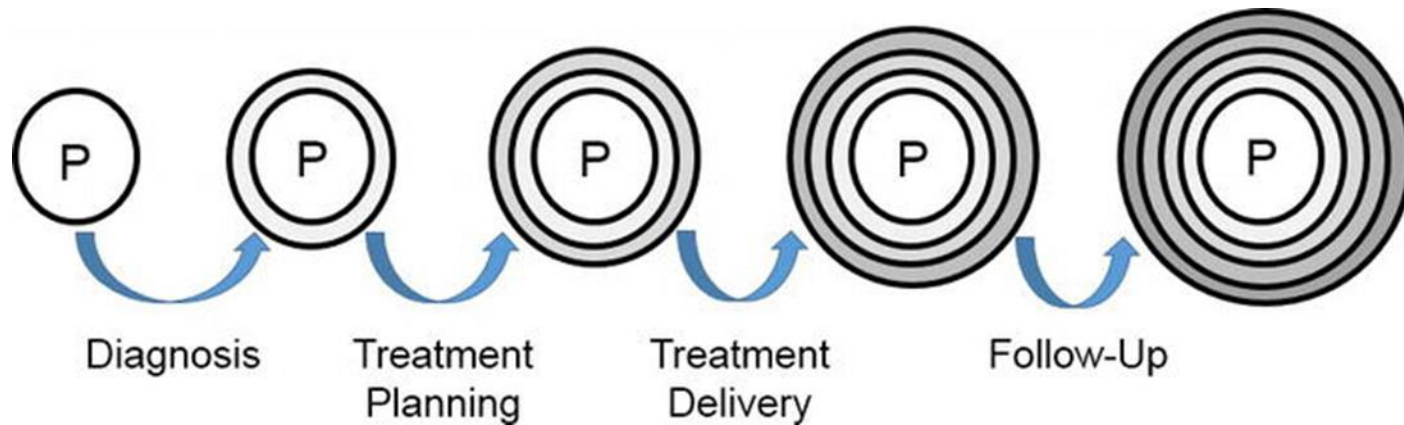
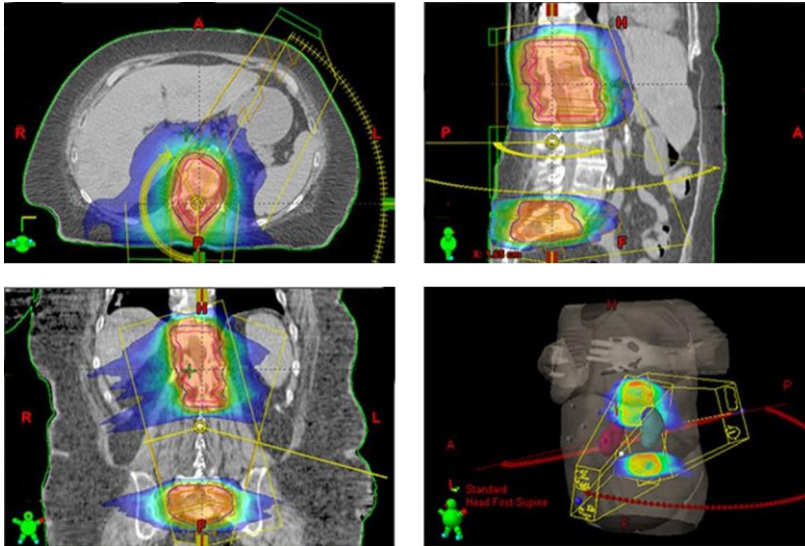


Figure 2: Overview of the proposed SCAN framework that jointly trains a segmentation network and a critic network with an adversarial mechanism. The segmentation network produces per-pixel class probabilities. The critic takes either the ground truth label or the prediction by the segmentation network, optionally with the CDR image, and output the probability estimate of whether the input is the ground truth (with training target 1) or the segmentation network prediction (with training target 0).

# Deep Reinforcement learning

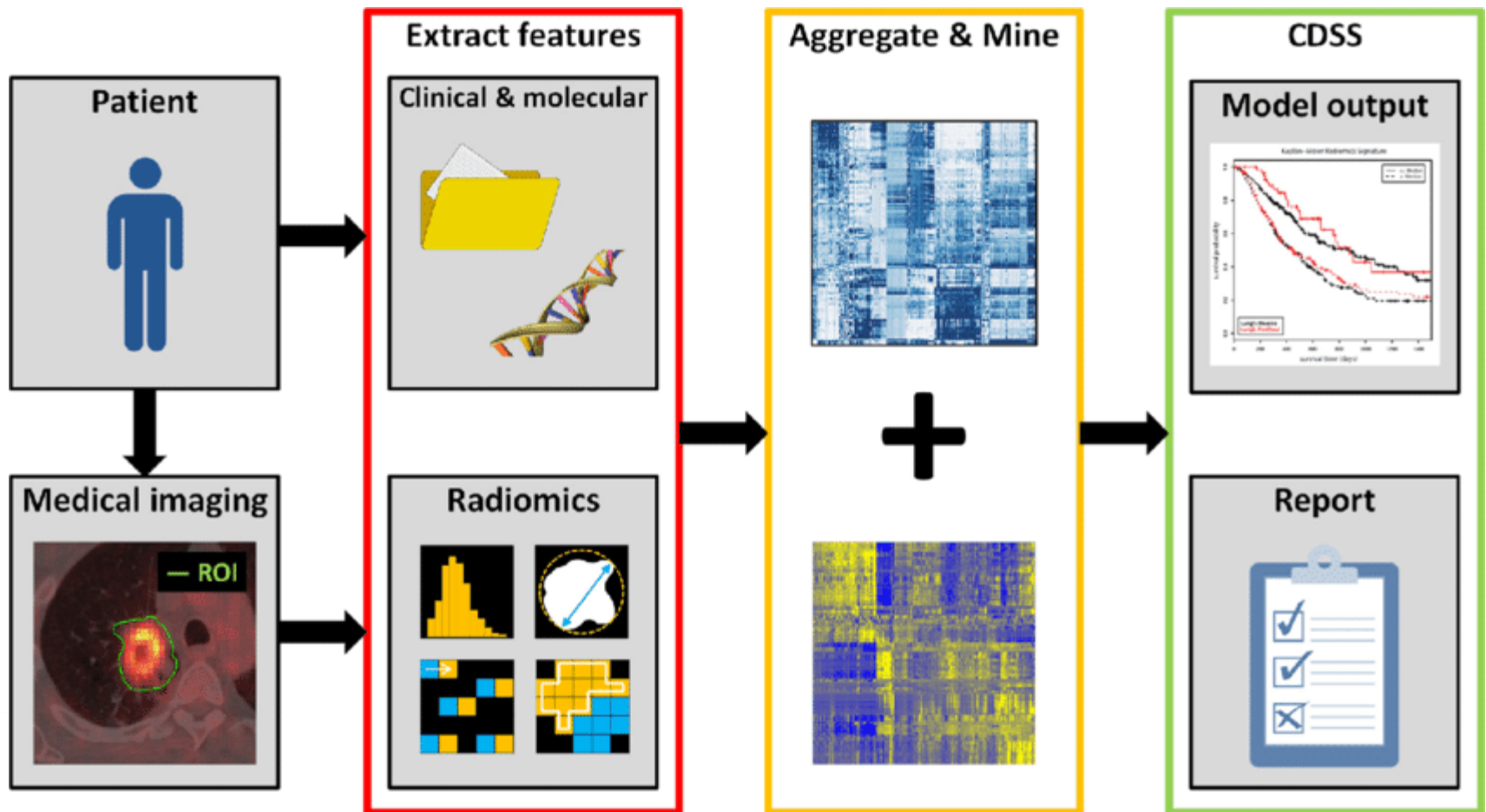


# Optimizing by Deep (R)L



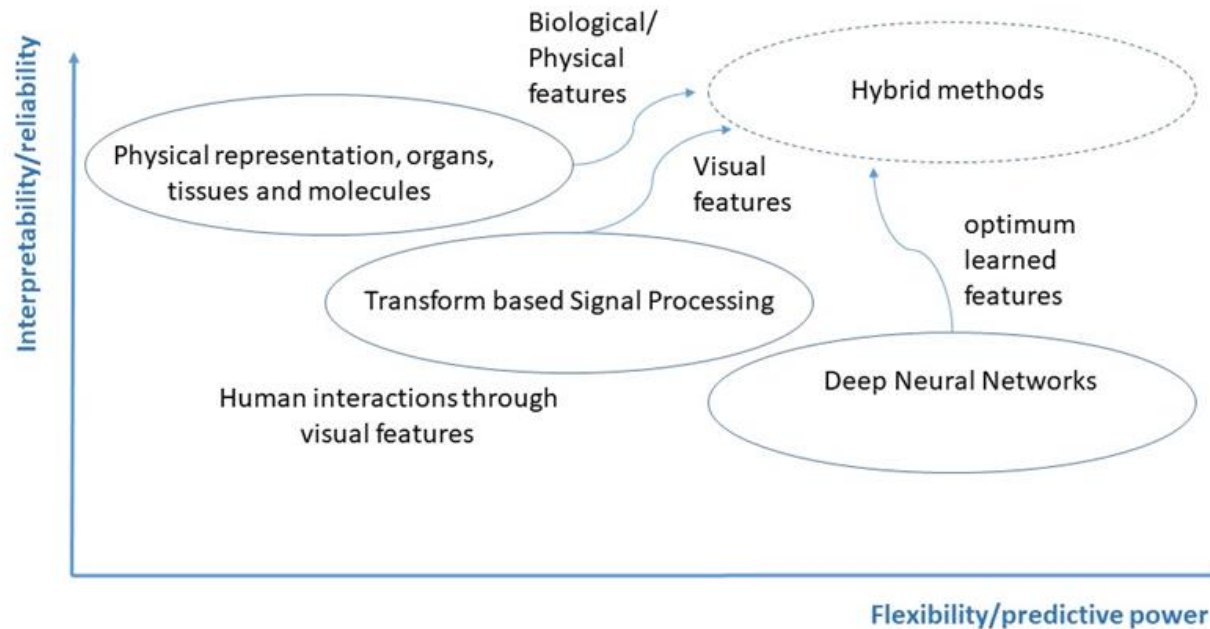
# The Radiomics challenge

- Predictive and personalized medicine



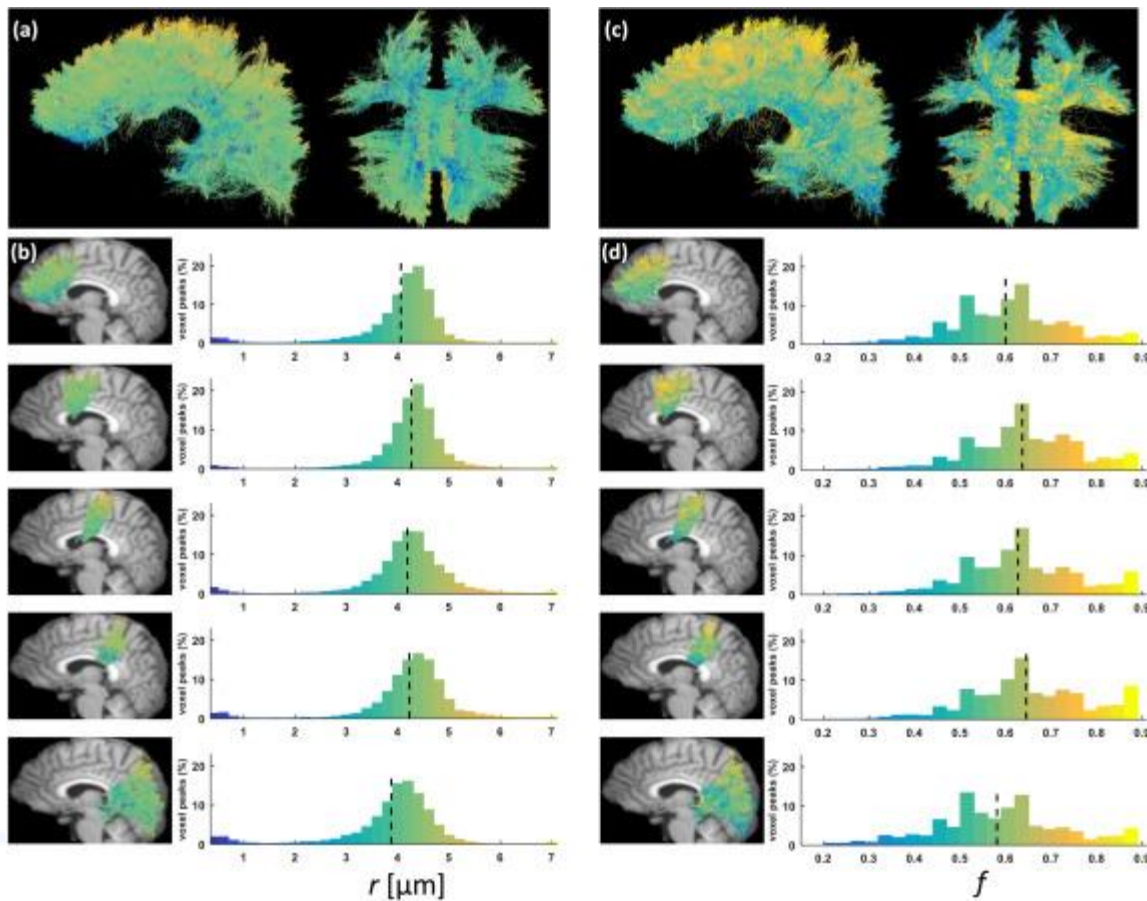


# Research assumption 1: three kinds of latent spaces by multi-agents



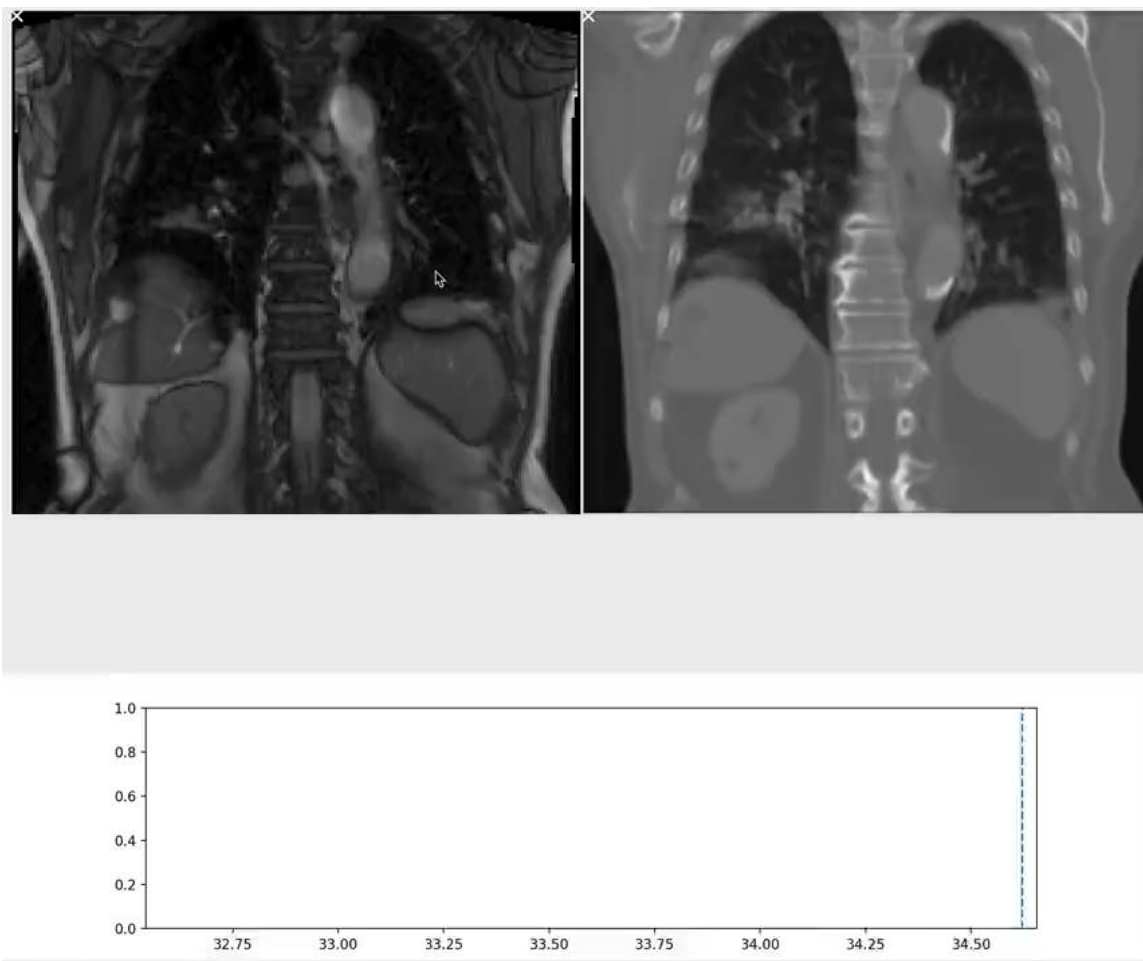
# Fingerprints (physical) latent space

- Rensonnet, G., Scherrer, B., Girard, G., Jankovski, A., Warfield, S. K., Macq, B., ... & Taquet, M. (2019). Towards microstructure fingerprinting: Estimation of tissue properties from a dictionary of Monte Carlo diffusion MRI simulations. *NeuroImage*, 184, 964-980.



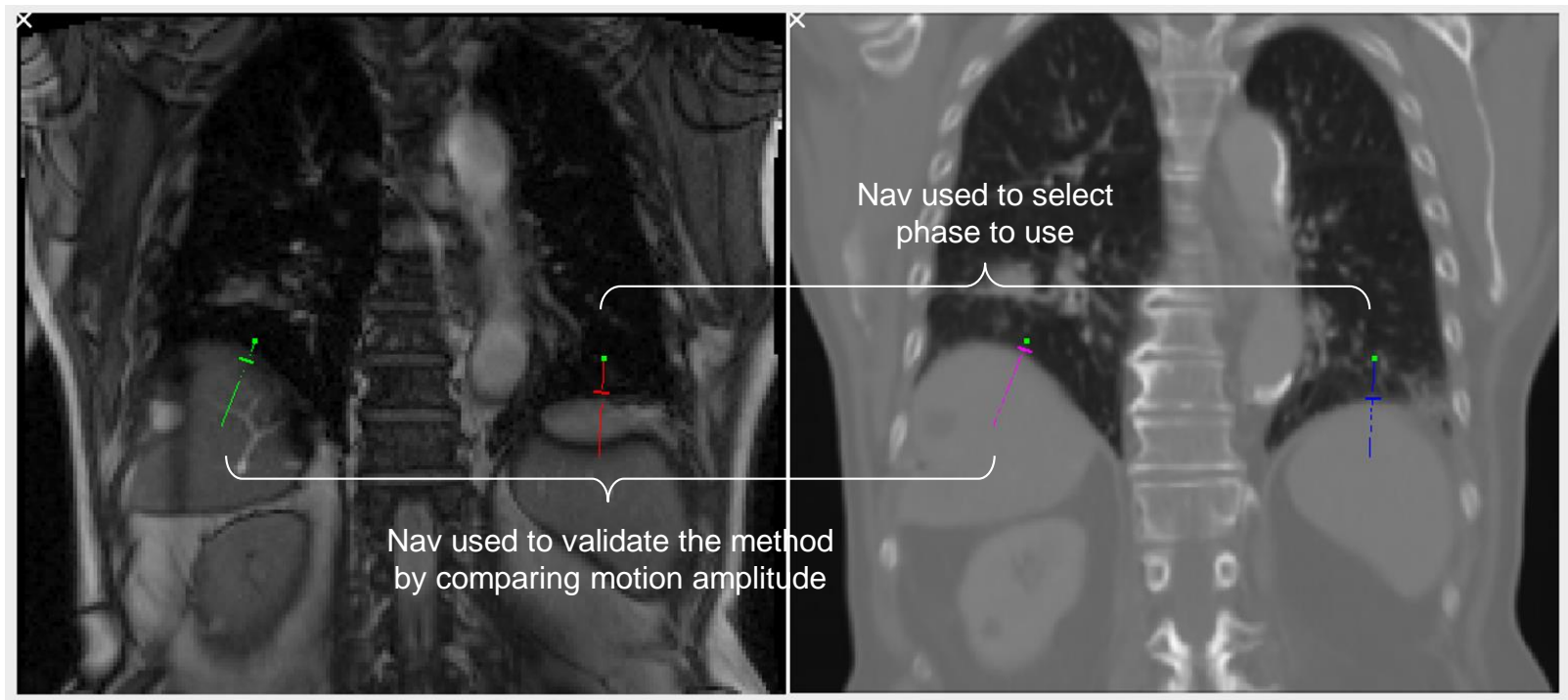
# Visual feature (actionable) latent space

Motion measurements 4DCT and IRM (coronal) after co-registration (2D on 3D)

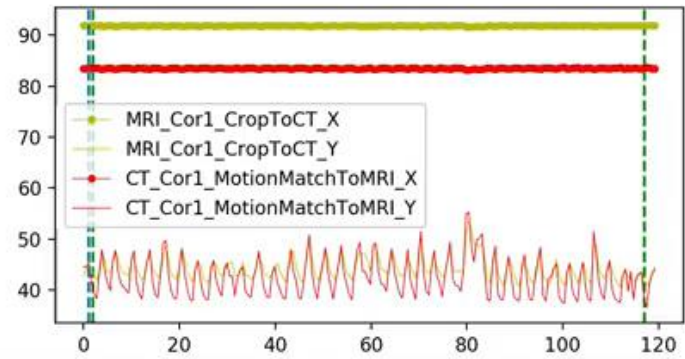
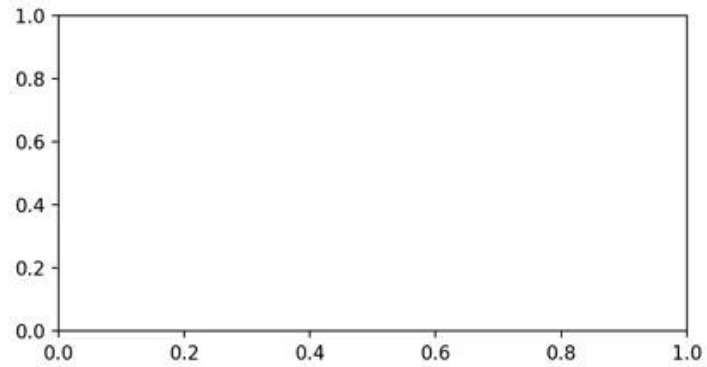
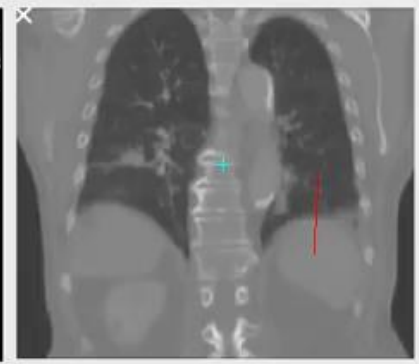
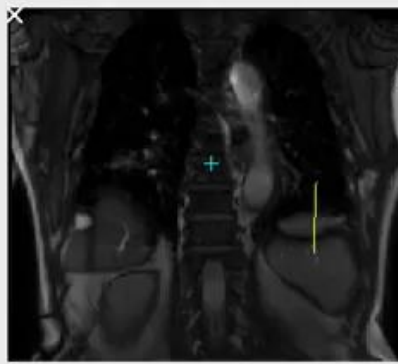
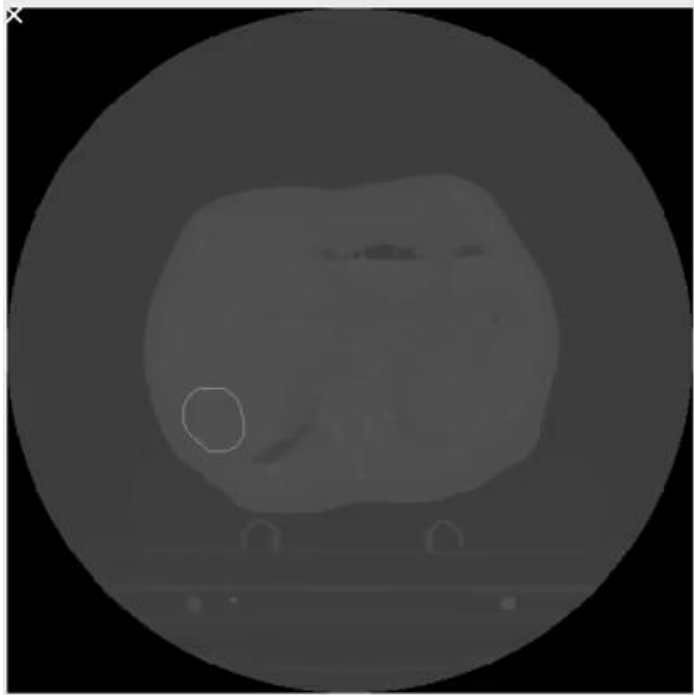


## Validation

- a. Comparison with motion at the same position
- b. Comparison with motions at other positions

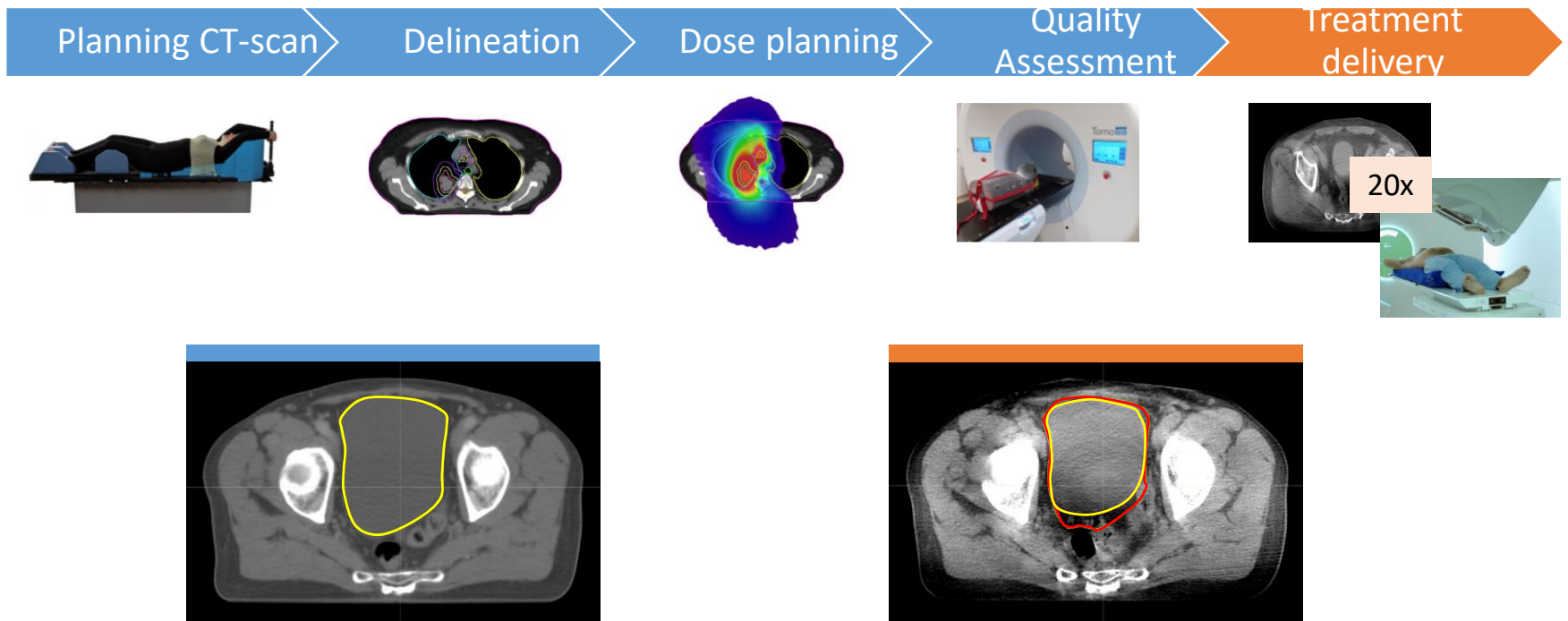


# Complete example with dose delivery observation

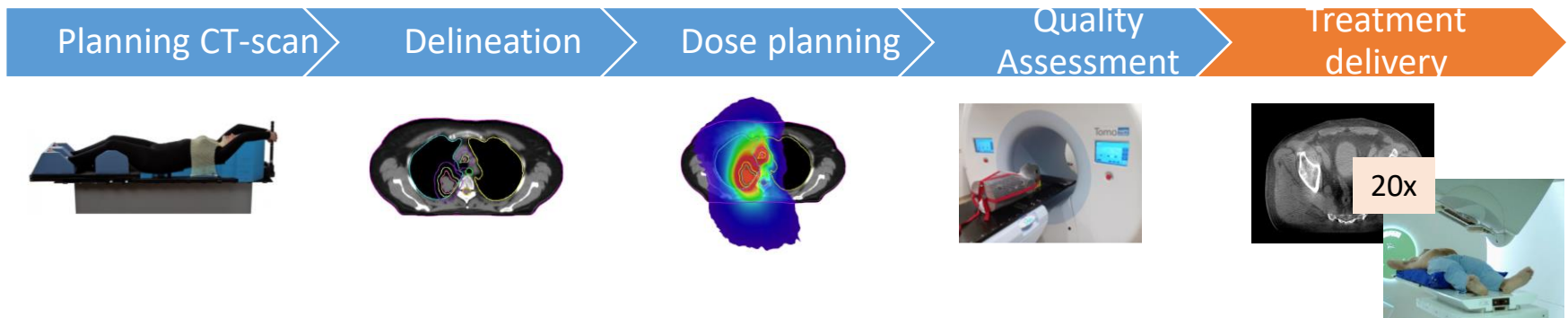


# Deep Learning latent space

Measuring anatomical variations between treatment sessions would improve dose conformity



# Measuring anatomical variations between treatment sessions would improve dose conformity



## Problem

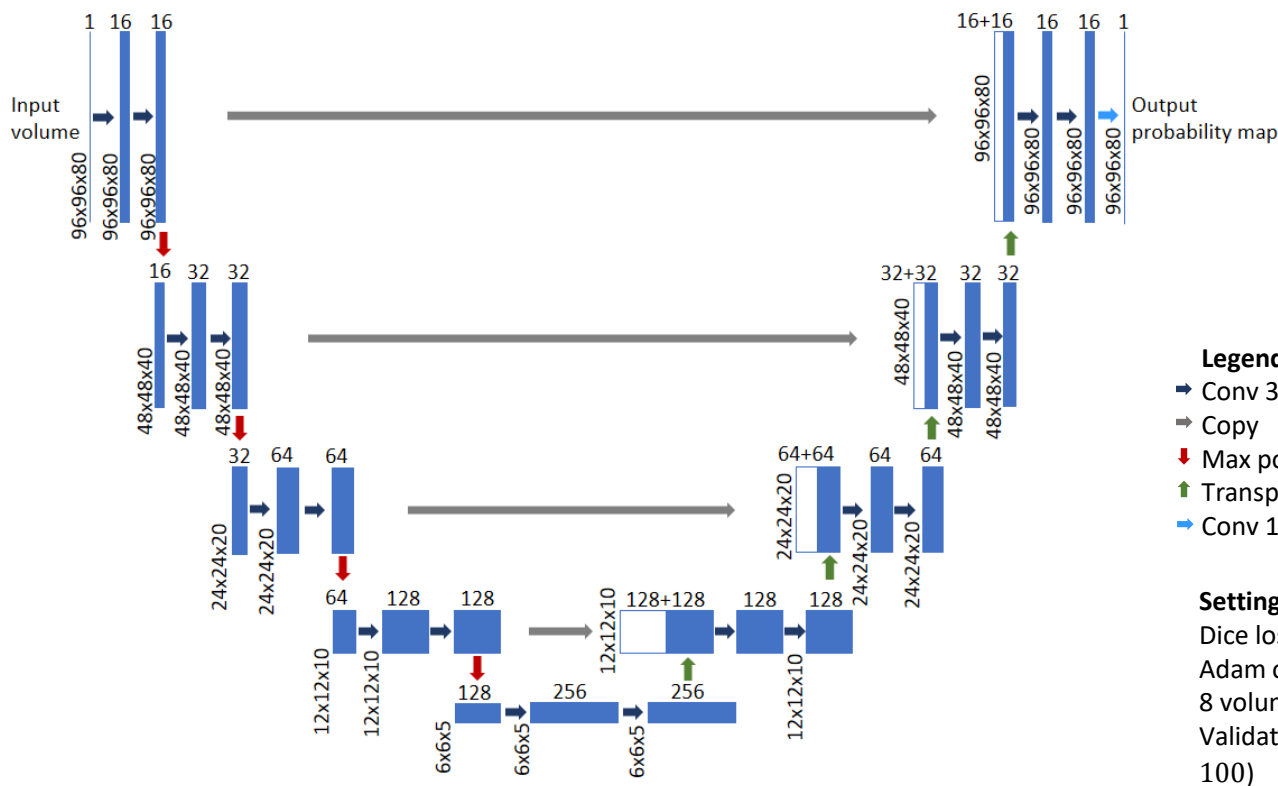
Scarcity of annotated CBCTs to train a deep neural network

## Question

Add (abundant) annotated CTs in training set?



# Methods: The network architecture is u-net



**Legend**

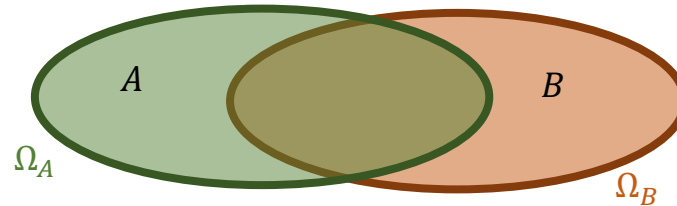
- Conv 3x3x3, ReLu, « same » padding
- Copy
- ↓ Max pooling 2x2x2
- ↑ Transpose conv 2x2x2, « same » padding
- Conv 1x1x1, Sigmoid

**Settings**

- Dice loss
- Adam optimizer ( $LR = 10^{-4}$ )
- 8 volumes per batch
- Validation set to earllystop training ( $n_{max\ epochs} = 100$ )
- Data augmentation (shift, rotation, shear)

Adapted from Ronneberger *et al.*, 2015

# Performance assessment



**Dice similarity coefficient**

$$\text{DSC} = \frac{2|A \cap B|}{|A| + |B|}$$

**Jaccard index**

$$\text{JI} = \frac{|A \cap B|}{|A \cup B|}$$

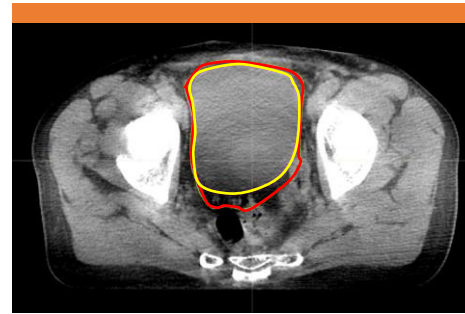
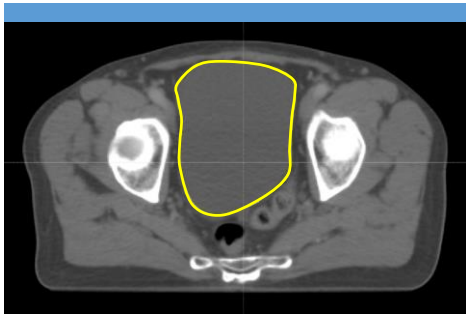
**Symmetric mean boundary distance**

$$\text{SMBD} = \frac{\bar{D}(A, B) + \bar{D}(B, A)}{2}$$

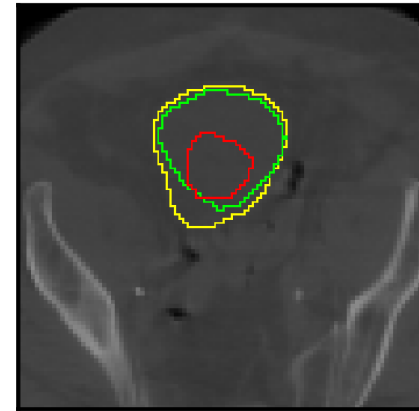
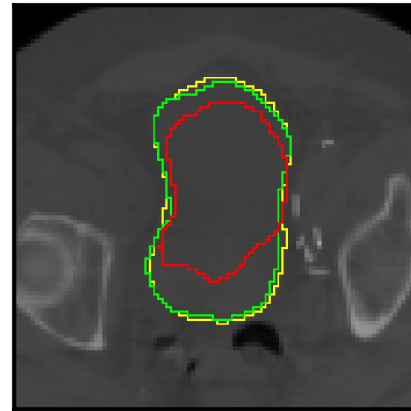
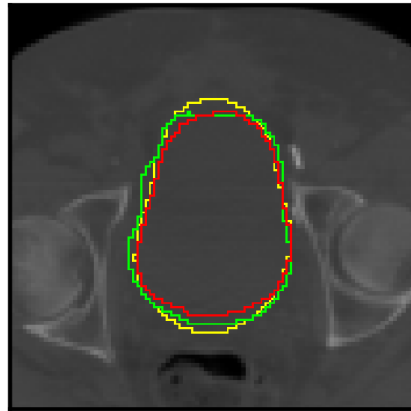
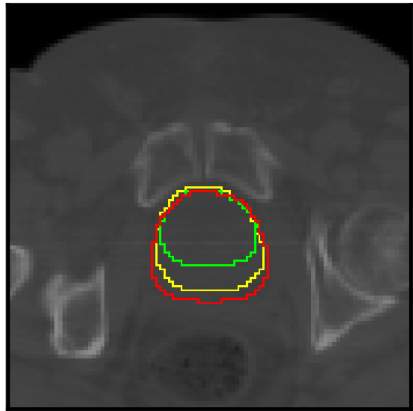
where  $D(A, B) = \left\{ \min_{x \in \Omega_B} \|x - y\|, y \in \Omega_A \right\}$

# Comparison baselines

## Deformable image registration



Results: Our approach outperforms a state-of-the-art DIR-based software on a representative patient

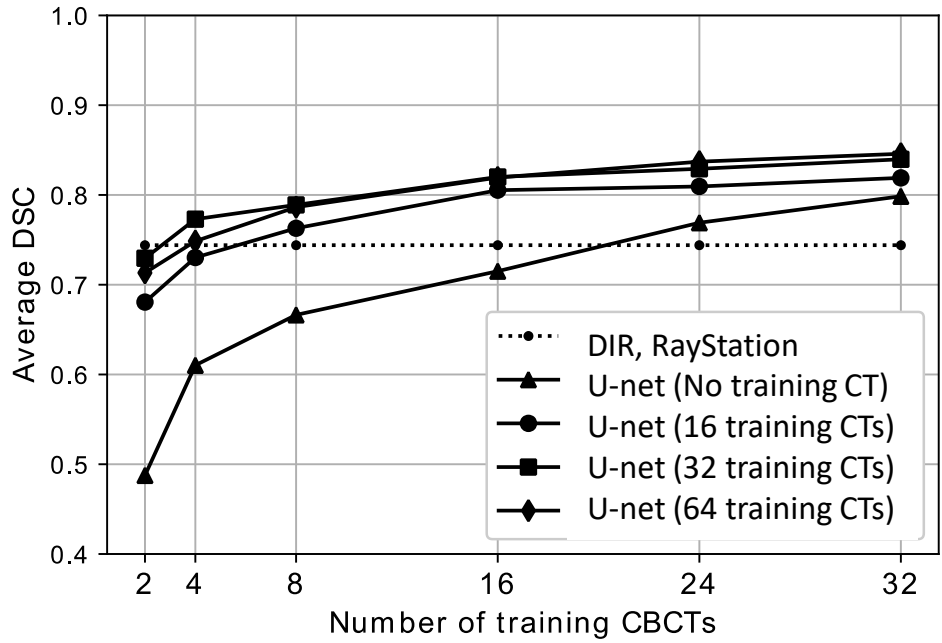


Ground truth segmentation

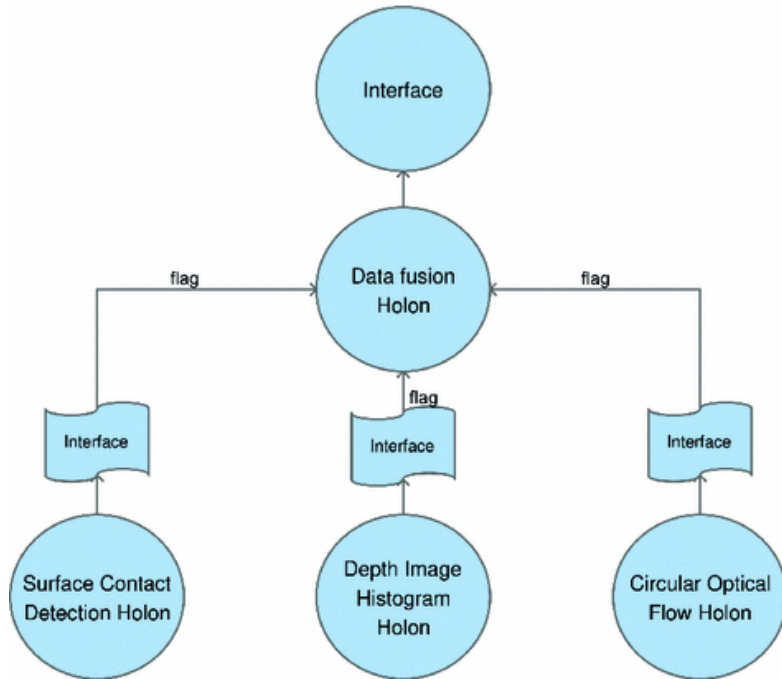
Deformable image registration, Raystation (DSC = 0.788)

U-net (DSC = 0.892, setting  $n_{CBCT} = 32$ ,  $n_{CT} = 64$ )

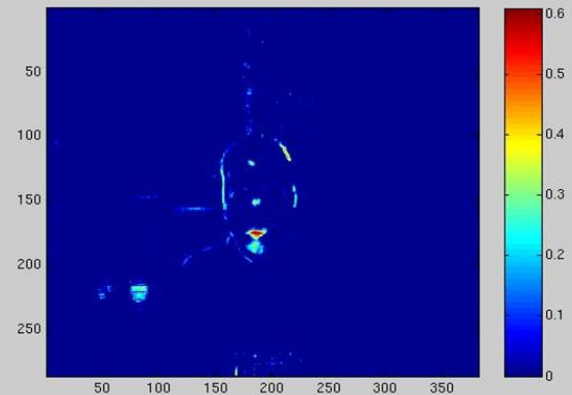
# Results



# 3 latent spaces cooperating in a multi-agent approach (incl HITL)



Mutual information between Pixels and audio



# Research assumption 2: Byzantine learning for sharing data and expertise

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- Need of integrative coalitions
  - To share data (privacy and relevance)
  - To explore complementarity, redundancy and equivalence of the algorithms
  - To assess co-evolution of algorithms and human expertise
  
  - By the use of consensus mechanisms  
(Federated Byzantine Agreements- blockchain)



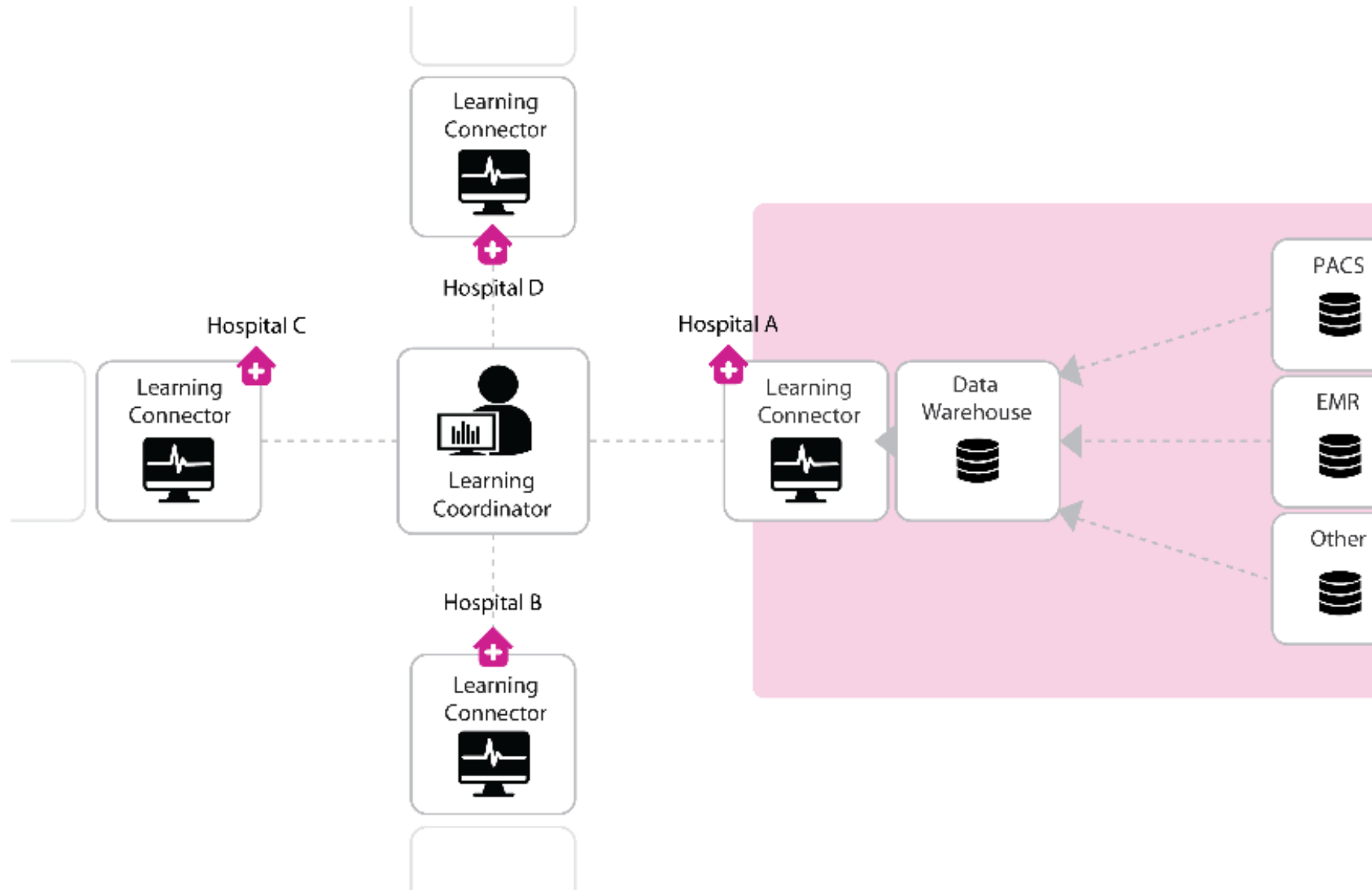
# The needs to better use Deep L

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## -Coalitions for Image Processing

- Distributed machine learning for larger data sets
- Trusted Image Processing through Integrative Coalitions
- Security (blockchains)
- Reliability (mutimodality-multiagents)
- Human in-the-loop (regular update-how to poll)

# The Oncoradiomics Model (Ph. Lambin)



# Distributed learning: an abundant literature

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- Distributed SVM: convergence equivalent to central learning can be proven

- Boyd**, Stephen, et al. "Distributed optimization and statistical learning via the alternating direction method of multipliers." Foundations and Trends® in Machine learning 3.1 (2010): 1-122

- Forero, P. A., Cano, A., & Giannakis, G. B. (2010). Consensus-based distributed support vector machines. Journal of Machine Learning Research, 11(May), 1663-1707.

- Distributed DNN – Federated learning convergence similar to central learning can be shown

- McMahan, B., & Ramage, D. (2017). Federated learning: Collaborative machine learning without centralized training data. Google Research Blog, 3.

# Security requirements

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## Challenge 1

**Data privacy** of the datasets used for the training (leakage effect of the gradients) : working by batches- differential privacy is the “crypto” model

## Challenge 2

**Protection** of the model against degradation by training on inadequate data: steps validation by the coalition and blockchained public ledger with hash of the iterative versions of the model

## Challenge 3

**Confidentiality** of the model and the gradients: homomorphic operations and/or access control of the model vault

## Challenge 4

**Traceability** of the model: DNN watermarking

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## WHERE MIGHT A DISTRIBUTED LEDGER USE CRYPTOGRAPHY?

### *Initiation and Broadcasting of Transaction*

- *Digital Signatures*
- *Private/Public Keys*

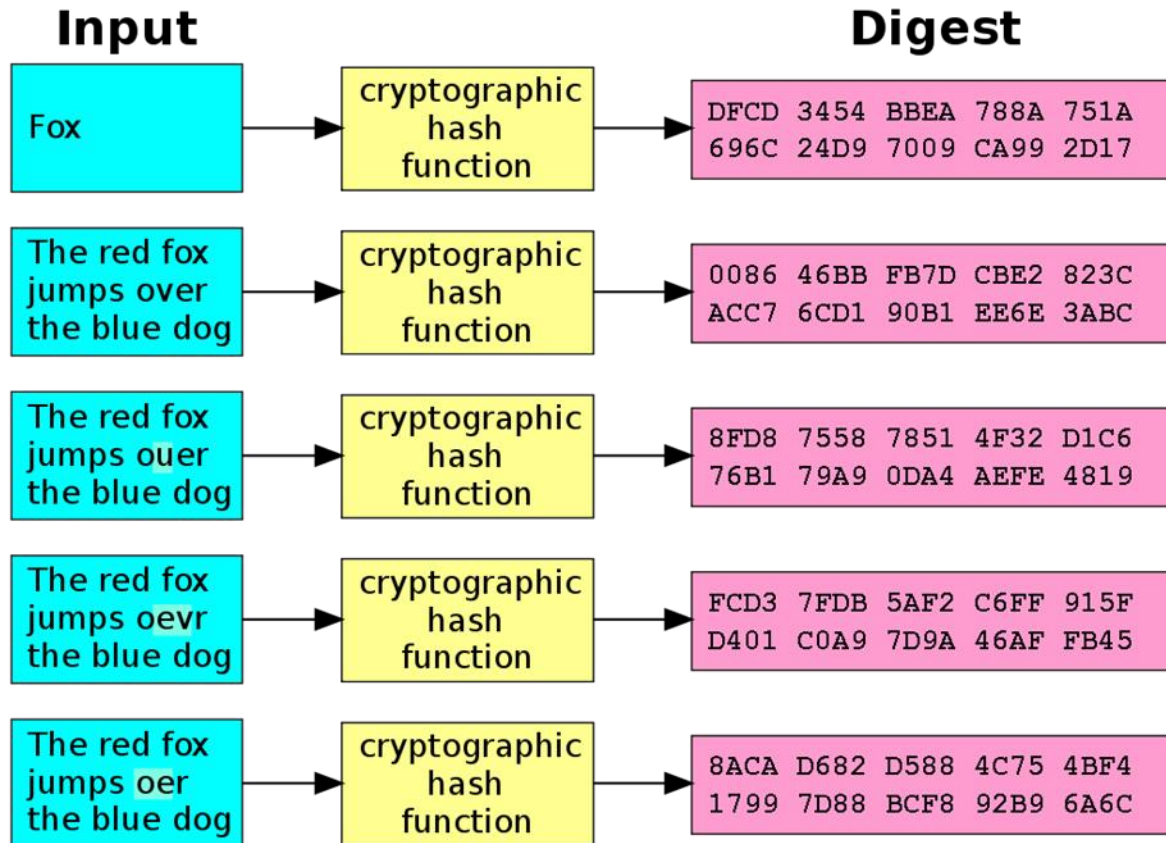
### *Validation of Transaction*

- *Proof of Work and certain alternatives*

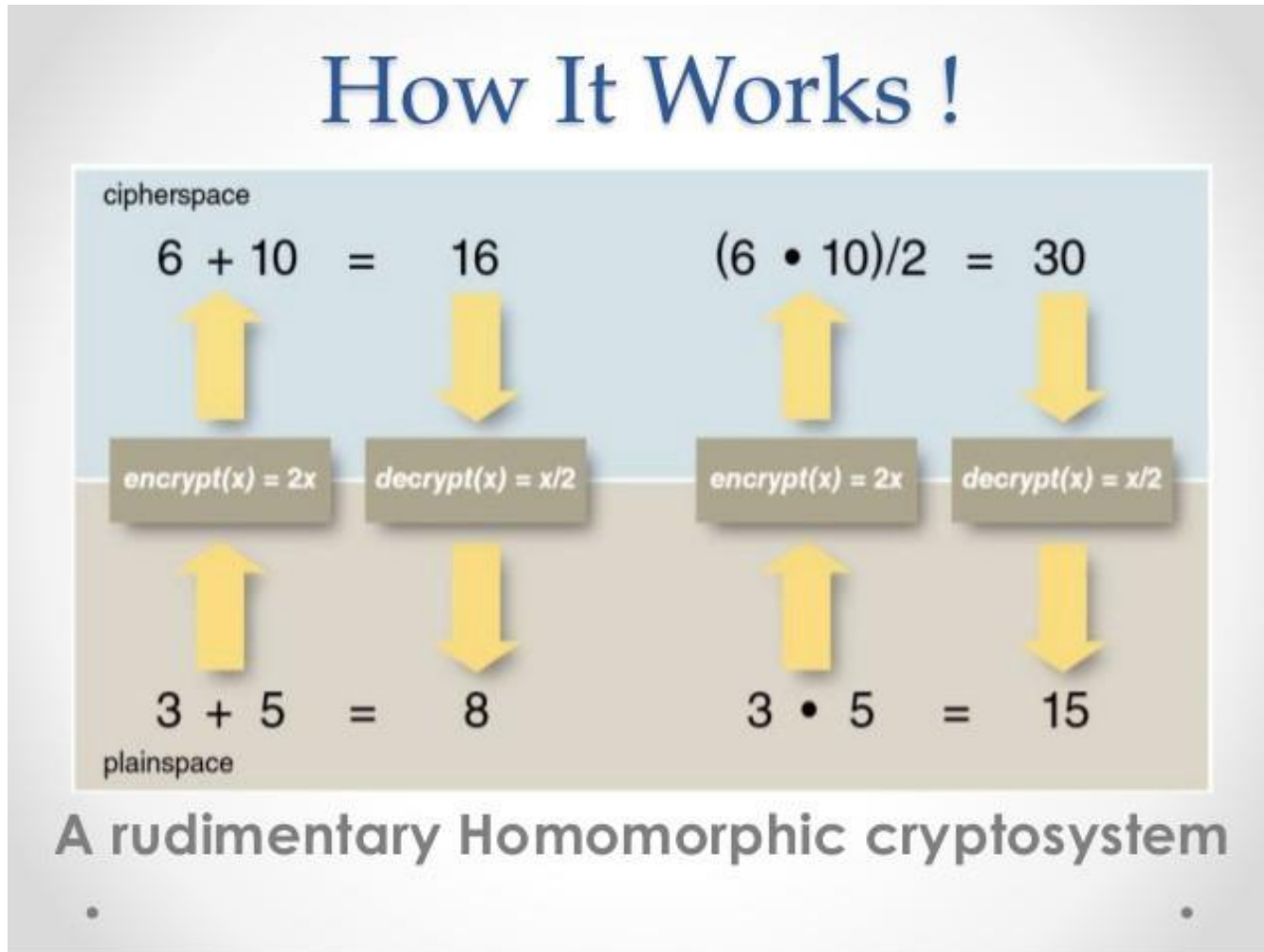
### *Chaining Blocks*

- *Hash Function*

# The hash function: SHA (one-way!!)



# Homomorphic encryption





# Watermarking

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Secret marks in audiovisual contents:

- Authentication
- Copyright
- Fingerprinting

Watermarks can be embedded into DNN:

-Uchida, Y., Nagai, Y., Sakazawa, S., & Satoh, S. I. (2017, June).

Embedding watermarks into deep neural networks. In Proceedings of the 2017 ACM on International Conference on Multimedia Retrieval (pp. 269-277). ACM.

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# Scalable security architectures for trusted coalitions

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## TCLearn-A

Learned model is *public*

Each member is accountable for the privacy protection of its own data

### **Solution to security challenge1**

(Data privacy of the datasets used for the training):

Local training of the model by each member with their own datasets

Generated gradients are uploaded and merged with the previous model

Batches of a minimum size to mitigate the long term memory effect

### **Solution to security challenge 2**

(Protection of the model against degradation by training on inadequate data):

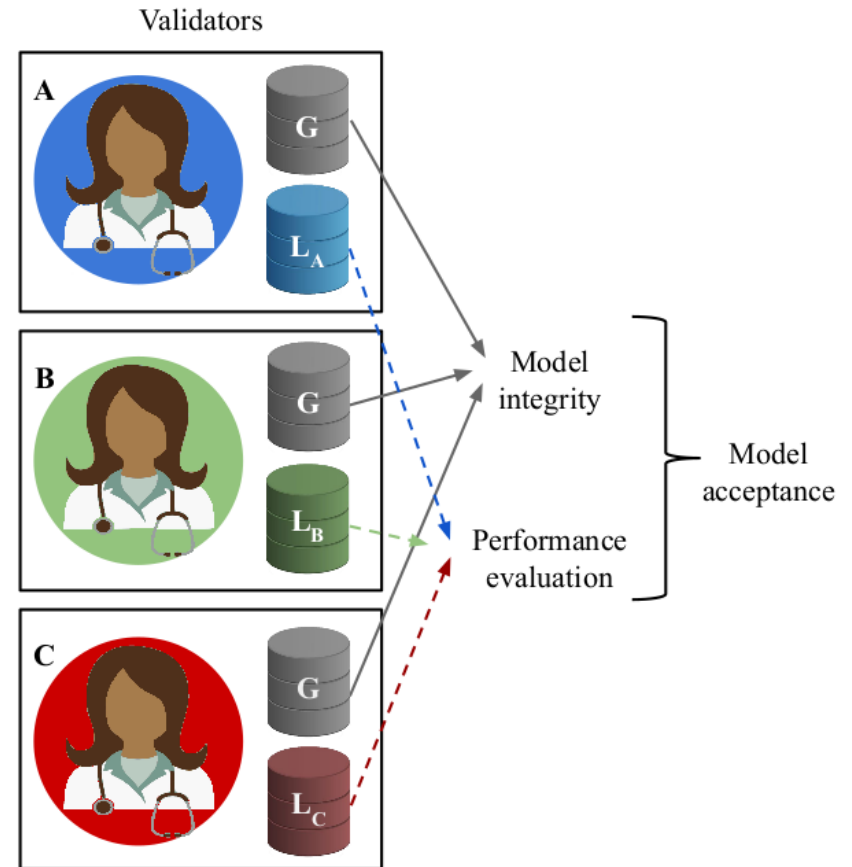
Blockchain storing cryptographic hashes of every training step

Federated Byzantine Agreement (FBA) to prevent corrupted increments

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# Federated Byzantine Agreement

- Two types of test databases: global test database (G), local test database (L)
- A “general” is randomly selected among the validators
- The “general” creates a new candidate block referencing the new model
- Every validator validates the viability (model) and integrity of this new candidate block
- Each validator broadcasts its opinion (positive or negative)
- The FBA process ends when 2/3 of the validators agree



# Scalable security architectures for trusted coalitions

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## TCLearn-B

Learned model is *private*, the members of the coalition trust each other.

### **Solution to security challenges 1 & 2:**

*Same as for TCLearn-A*

### **Solution to security challenge 3:**

(Confidentiality of the model and the gradients):

Storage of all iterations of the model in an off-chain storage

Iterations only referenced by links in the blockchain

Secure, encrypted transport of the model (using e.g. TLS or S/MIME)

### **Solution to security challenge 4:**

(Traceability of the model):

Access control and audit mechanisms to protect the models and parameters

# Scalable security architectures for trusted coalitions

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## TCLearn-C

The members of the coalition do not trust each other.

### **Solution to security challenges 1 & 2:**

*Same as for TCLearn-A*

### **Solution to security challenges 3 & 4:**

Storage of all iterations of the model in an off-chain storage

Each member is provided with a homomorphically encrypted model and the corresponding public key, used to encrypt their datasets, by a supervisor

Prediction could be performed locally on encrypted data, but the result must be decrypted by the supervisor

Full traceability since the encrypted model cannot be used without the associated public key, itself associated with the partner which received it

# Summary of our blockchain D DNN

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- New architecture for distributed learning based on a blockchain using a federated Byzantine agreement
  - Performance of the model ensured through shared evaluation of individual contributions (leading to acceptance or rejection)
  - Trusted coalitions, actions for updating the model stored on a public ledger implemented as a blockchain
  - Three kinds of coalitions with increasing security levels depending on the requirements for the distribution of the model
  - Solutions based on effective cryptographic tools and homographic encryption
  - Data privacy protection through encryption and off-chain storage
  - <https://arxiv.org/abs/1906.07690> (Lugan .... Macq)
-