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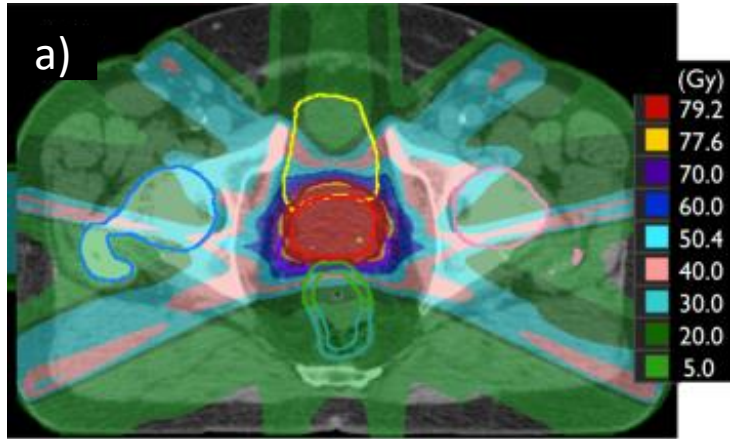


Proton Range Verification using Protoacoustics and Artificial Intelligence

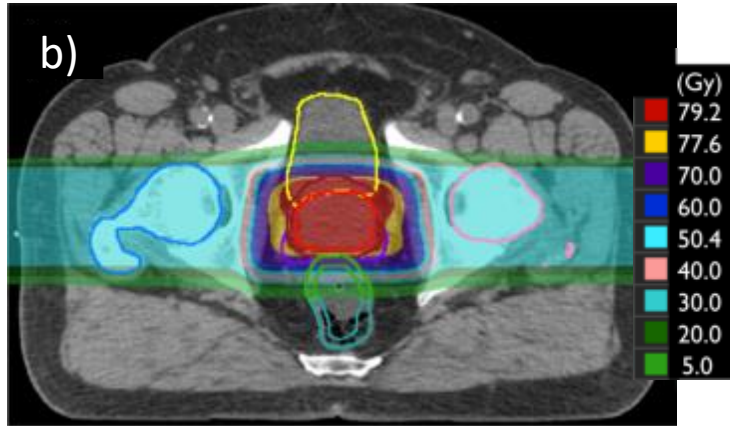
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Grupo de Física Nuclear, Dpto EMFTEL & IPARCOS, Universidad Complutense de Madrid

Instituto de Investigación Sanitaria del Hospital Clínico San Carlos (IdISSC)



Photon
Dose
Distribution

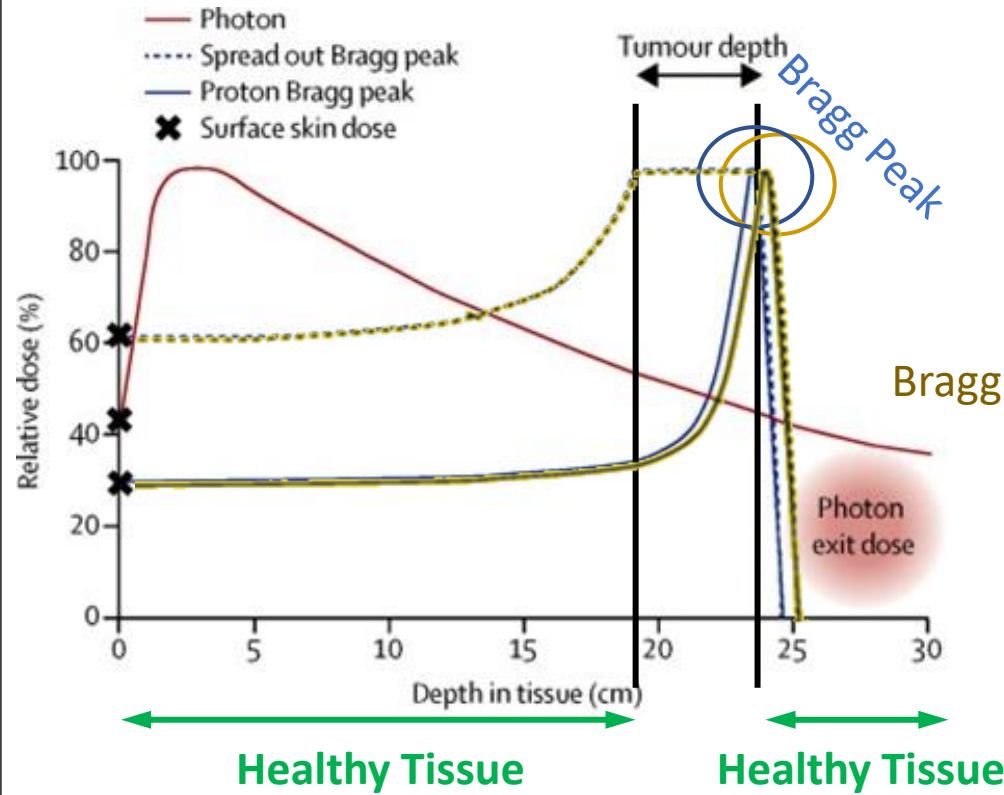


Proton
Dose
Distribution

Benefits:

- More conformal dose distribution
- Lower dose in healthy tissue

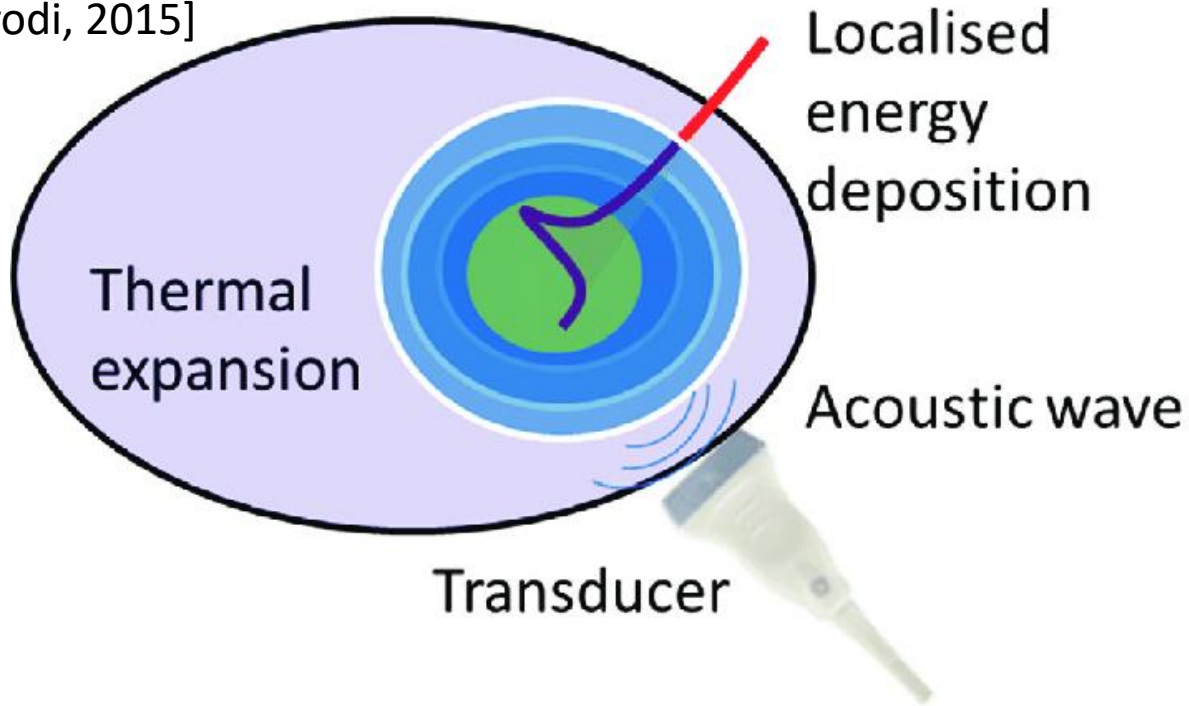
Problem: Proton Range Uncertainty



Solution:
Proton Range
Verification

Protoacoustic. Thermoacoustic effect

[Parodi, 2015]



Initial pressure distribution

Dose Density

$$p_0(\vec{r}) = \Gamma D(\vec{r}) \rho(\vec{r})$$

$$\Gamma = \frac{\beta c^2}{C_p}$$

Grüneisen coefficient

$$\Gamma_{water} = 0.11$$

$$p_0(\vec{r}, t) = p_0(\vec{r})G(t)$$

proton pulse temporal profile

Wave propagation:

$$\left(\nabla^2 - \frac{1}{c^2} \frac{\partial^2}{\partial t^2} \right) p(\mathbf{r}, t) = -\Gamma \frac{\partial}{\partial t} \mathcal{H}(\mathbf{r}, t)$$

Wave equation

Source term: deposited energy

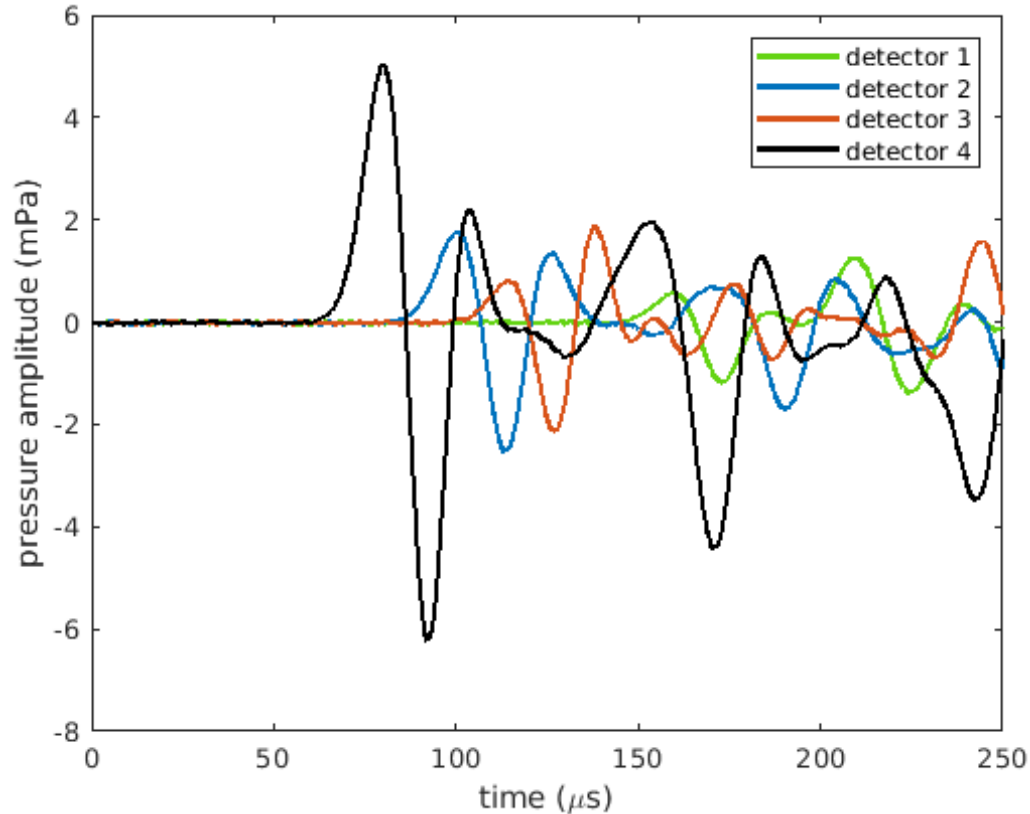
Solution (in homogeneous media):

$$p(\mathbf{r}_s, t) = \frac{1}{4\pi c} \frac{\partial}{\partial t} \int_{|\mathbf{r}_s - \mathbf{r}| = ct} \frac{p_0(\mathbf{r})}{|\mathbf{r}_s - \mathbf{r}|} \delta(t) d\mathbf{r}$$

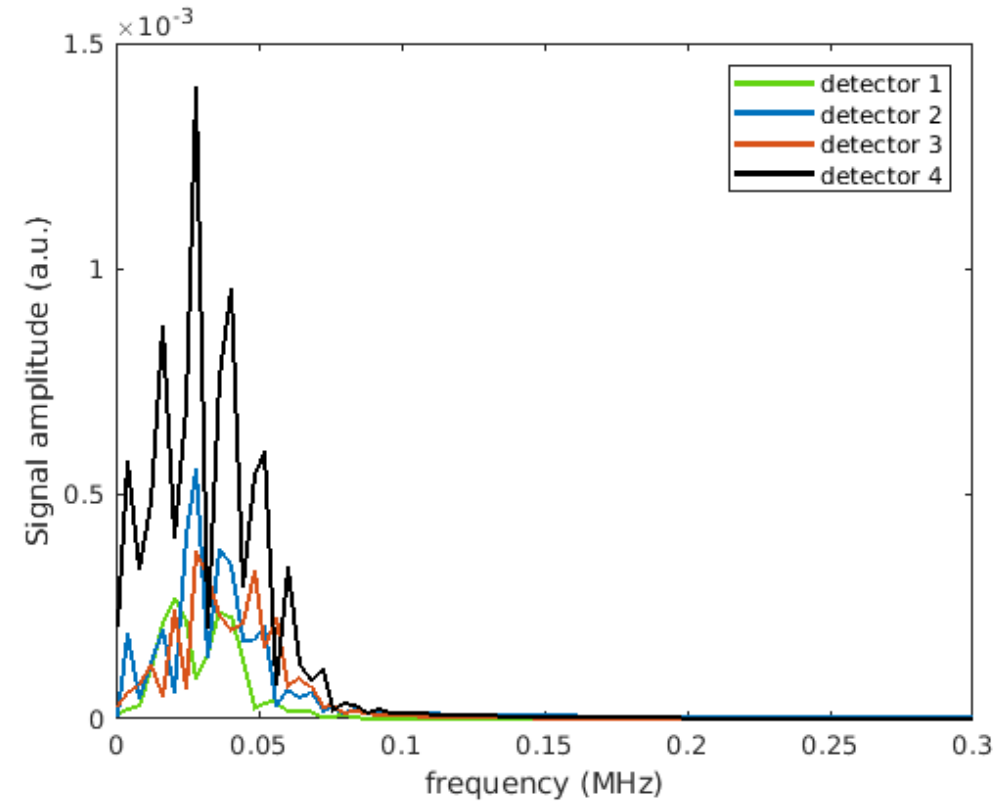
Time dependence Spatial dependence

Example of Protoacoustic Signals

Simulated Acoustic Signals induced by Proton Beam

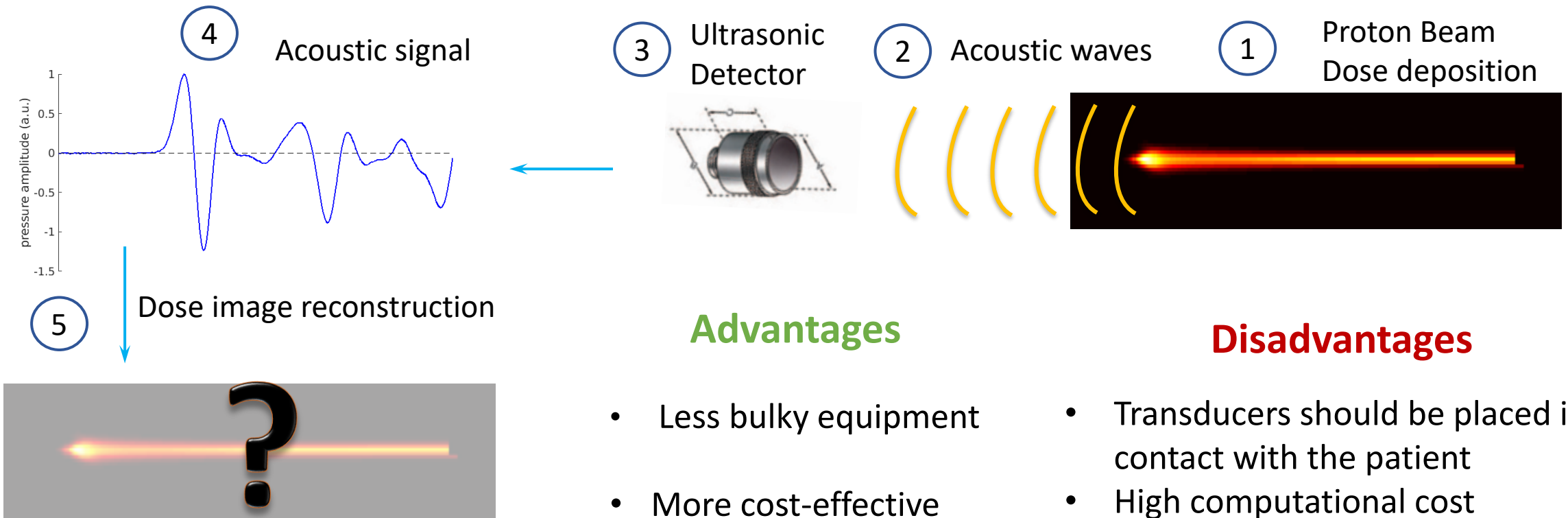


Maximum pressure
amplitude \sim mPa



- Frequency spectrum from **8 to 100 kHz**
- Central frequency of **28 kHz**
- Frequency \ll 1 MHz \rightarrow
Attenuation negligible in soft tissue

- **Nuclear activation techniques: PET and prompt-gamma imaging**
- **Alternative technique: Protoacoustics**

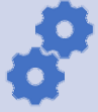


Advantages

- Less bulky equipment
- More cost-effective

Disadvantages

- Transducers should be placed in contact with the patient
- High computational cost



METHOD 1 – Iterative Reconstruction



METHOD 2 – Dictionary-based Reconstruction



METHOD 3 – Deep-Learning-based Reconstruction



1) Dose Reconstruction with a Linear Model

Measured
Pressure (y)

Model Matrix (A)

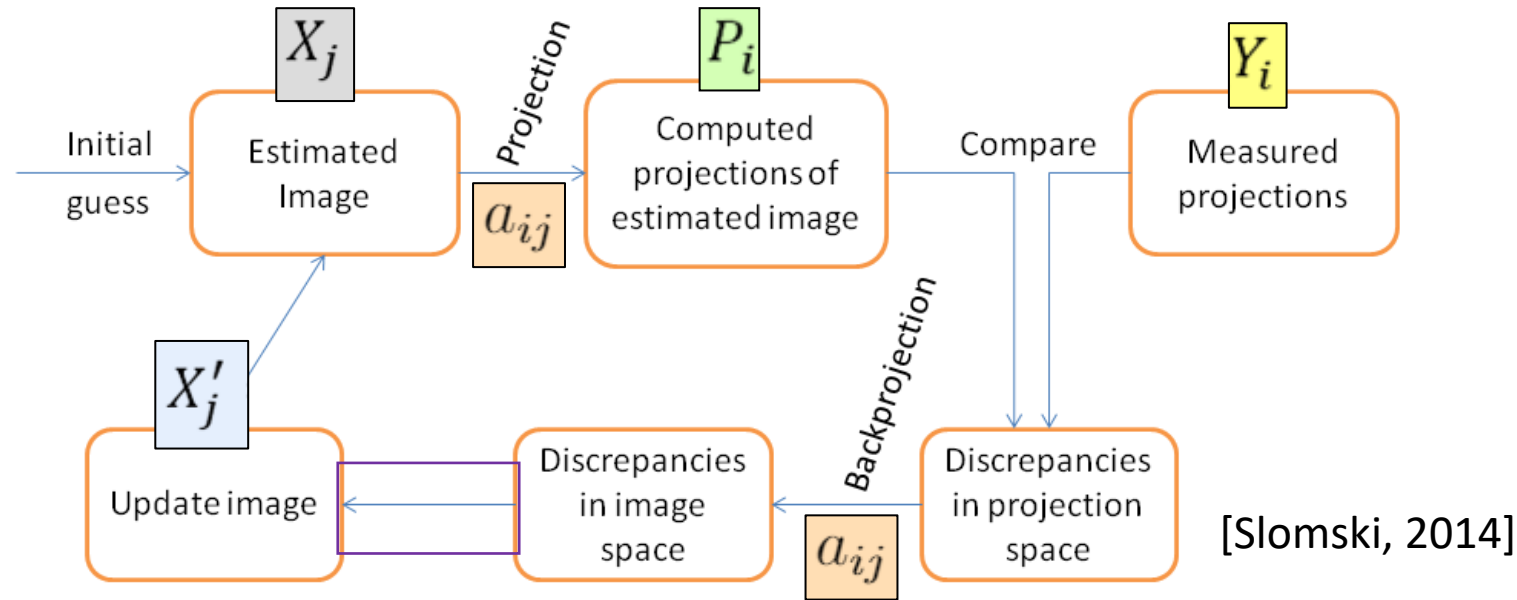
Initial
Pressure (x)

$$p_0(\vec{r}) = \Gamma D(\vec{r}) \rho(\vec{r})$$

$$p(\vec{r}_s, t) = \frac{1}{4\pi c} \frac{\partial}{\partial t} \int_{|\vec{r}_s - \vec{r}| = ct} \frac{p_0(\vec{r})}{|\vec{r}_s - \vec{r}|} \delta(t) d\vec{r}$$

$$y = Ax$$

1) Iterative Reconstruction Algorithms



Gradient-descent algorithm

$$X'_j = X_j + 2\lambda \sum_i (Y_i - P_i) \cdot a_{ij}$$

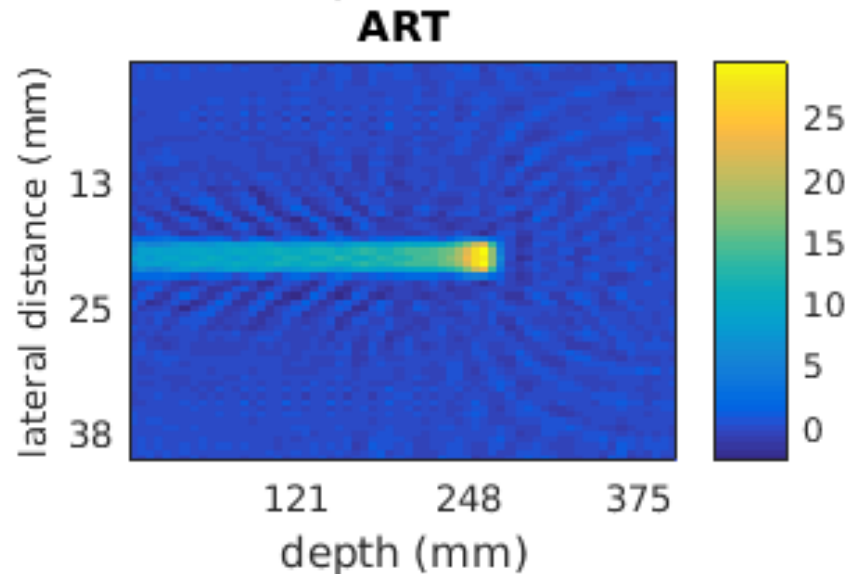
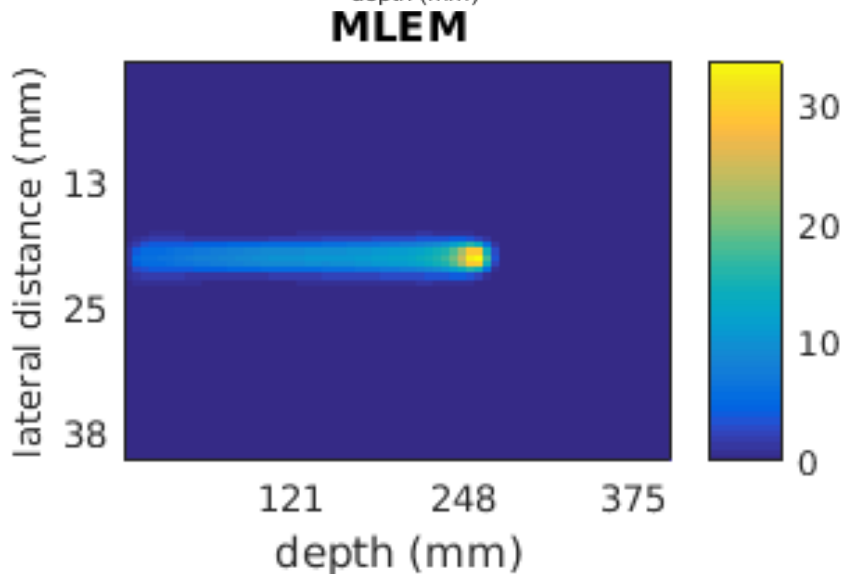
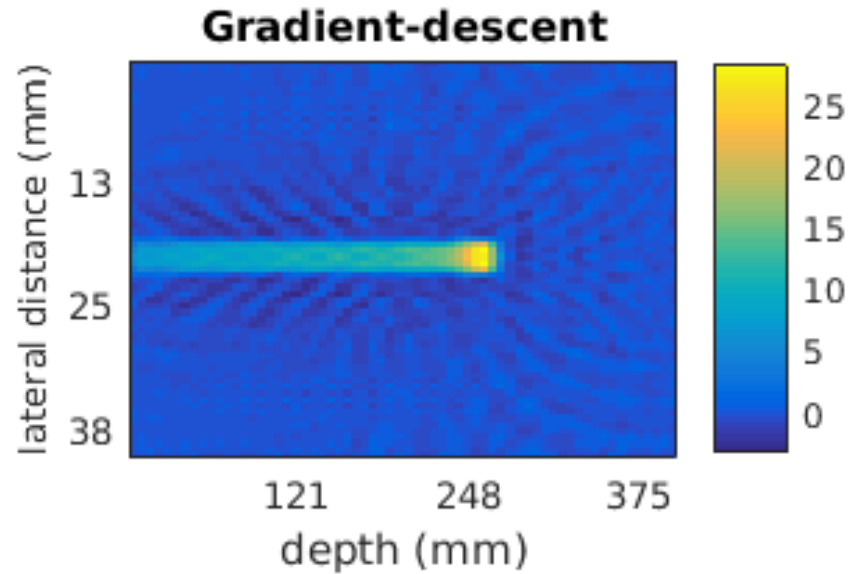
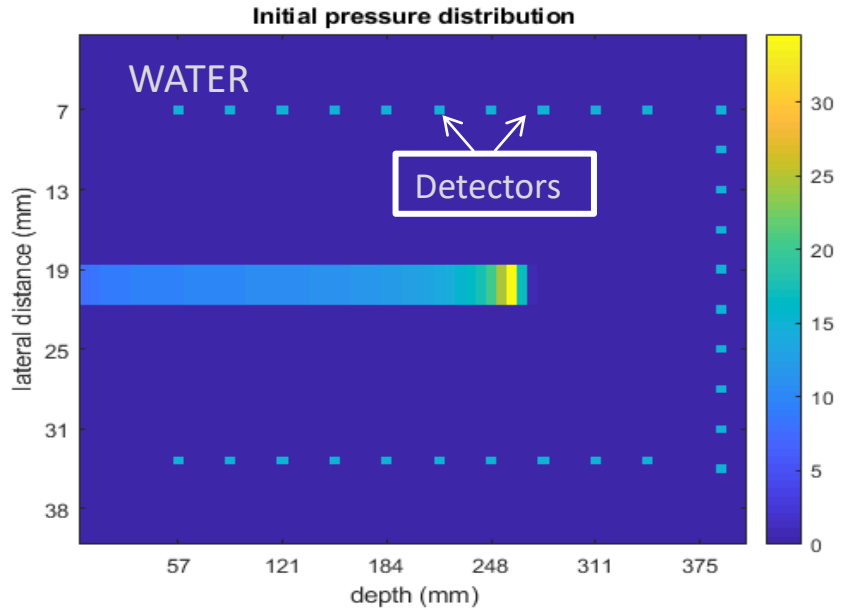
ART algorithm

$$X'_j = X_j + \frac{\sum_i (Y_i - P_i) \cdot a_{ij}}{\sum_i a_{ij}^2}$$

Modified MLEM algorithm

$$X'_j = X_j \cdot \frac{\sum_i (\hat{Y}_i / \hat{P}_i) \cdot \hat{a}_{ij}}{\sum_i \hat{a}_{ij}}$$

1) Example: Proton Dose in Water Simulation



Acoustic Simulation



k-Wave
A MATLAB toolbox for the time-domain
simulation of acoustic wave fields

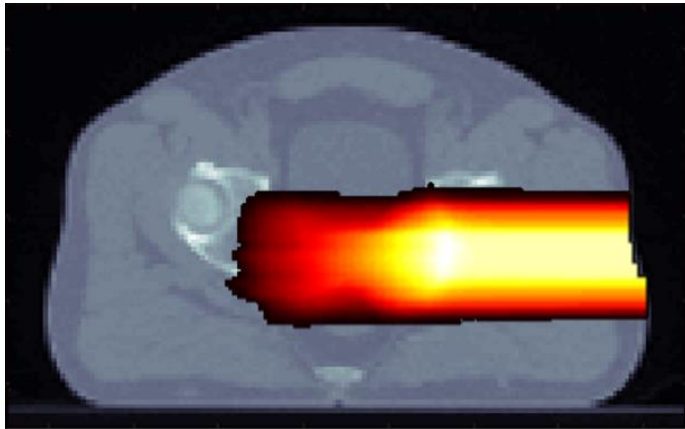
<http://www.k-wave.org/>

200 MeV
protons
In water

Method 2: Protoacoustic Dictionary

Workflow for Dictionary Construction

Treatment Plan



- Thousands of pencil beams
- Personalized for each patient
- Available at least one day before the treatment session

A priori Information

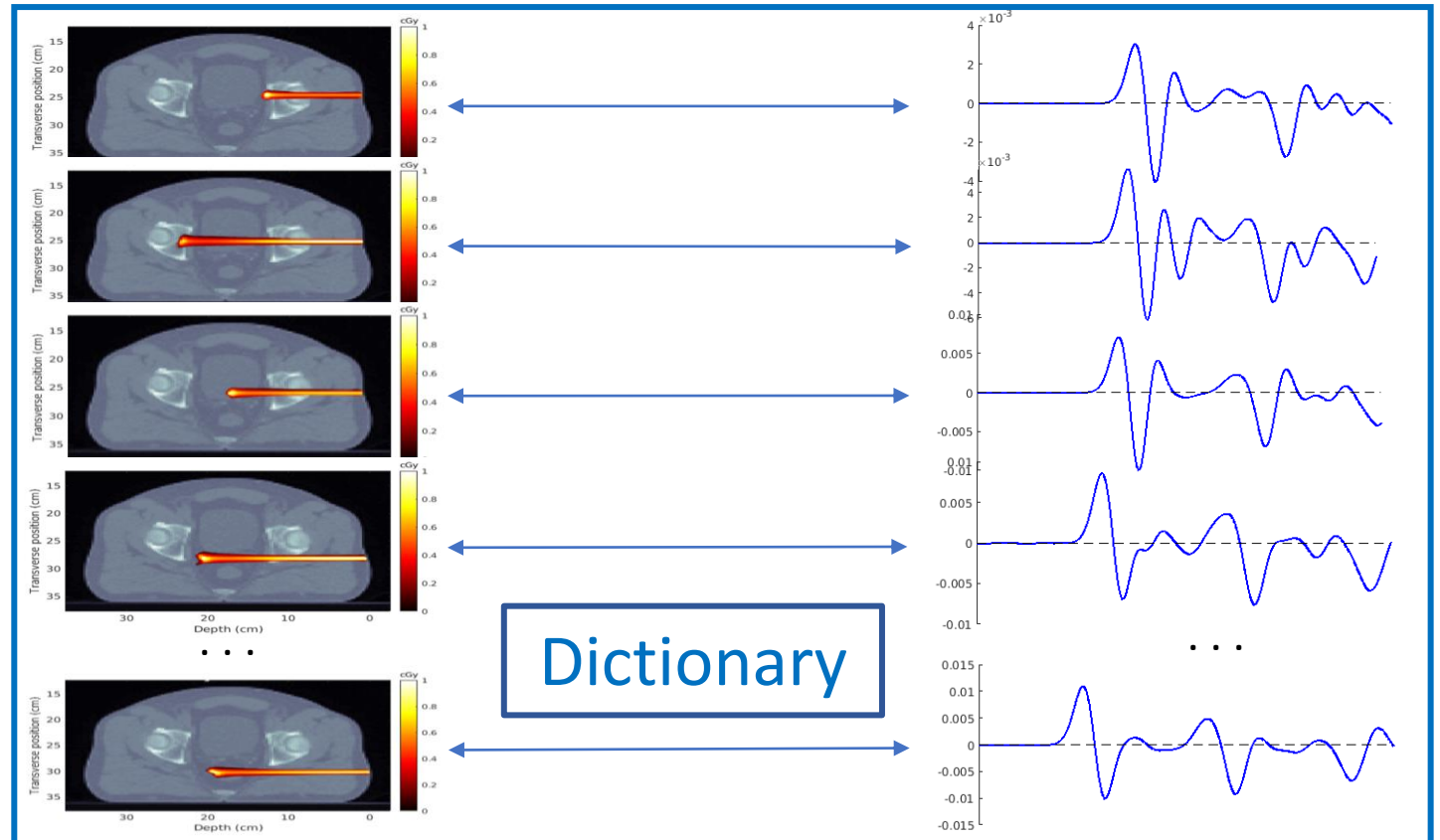
- Energy and pencil beam position
- Density
- Speed of sound

Acoustic Simulation



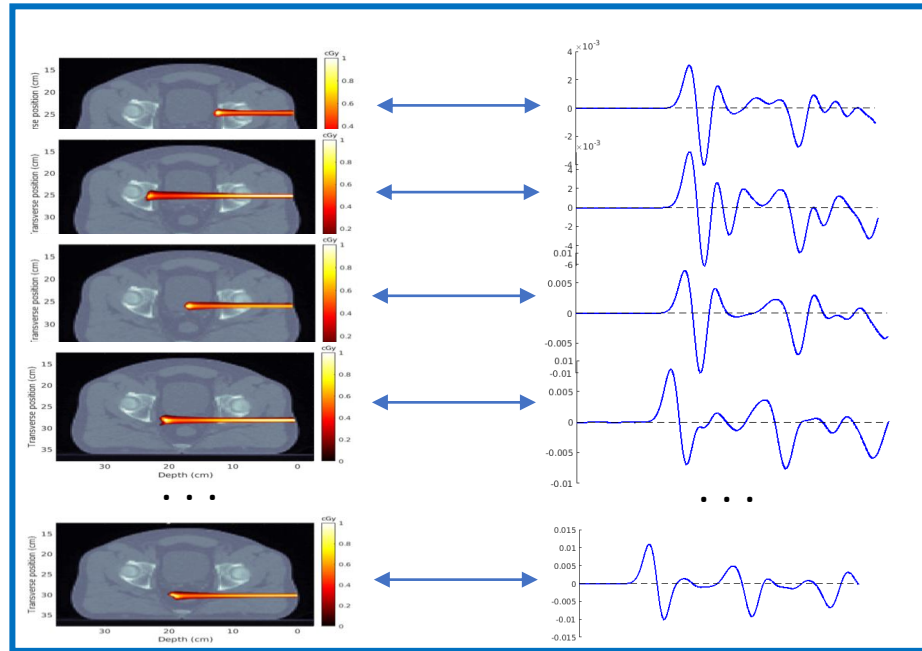
k-Wave
A MATLAB toolbox for the time-domain simulation of acoustic wave fields

<http://www.k-wave.org/>



2) Dictionary-based Proton Range Verification

Pre-Computation

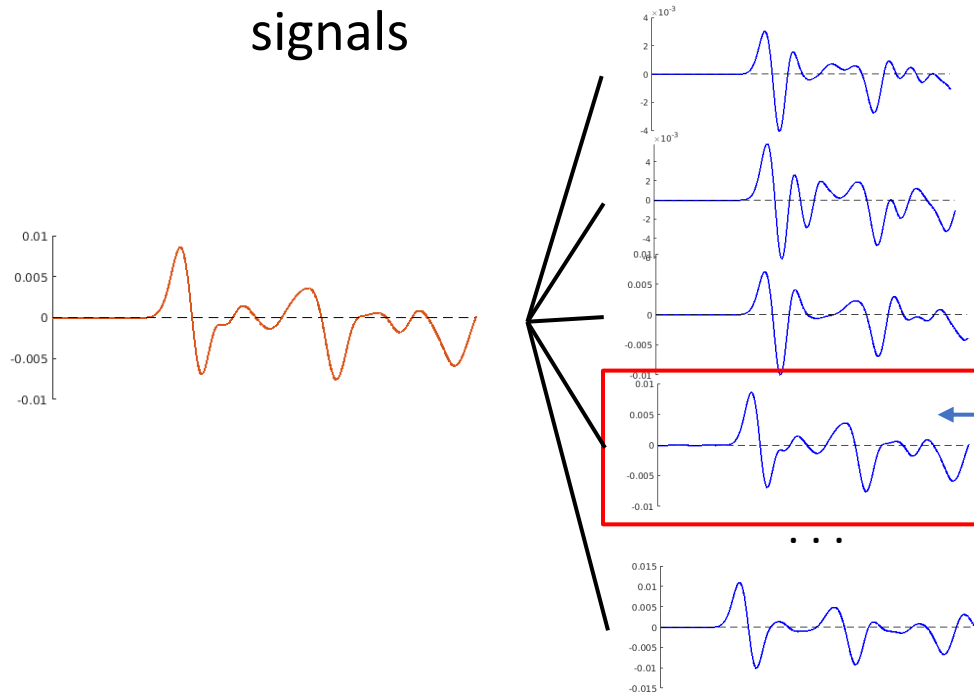


Individualized Dictionary

500 pencil beams ~ 100 min

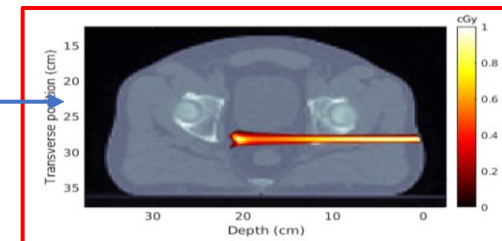
Real Time

Similarity
Measured vs Dictionary
signals



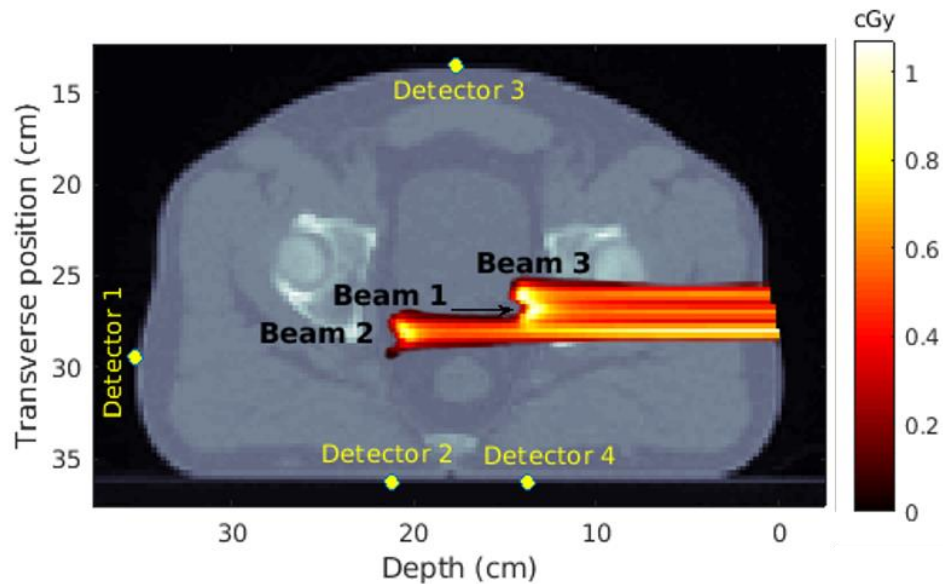
In-house developed
algorithm based on
L1 norm (100 ms)

Most probable
Bragg peak (BP)
position



2) Detection of Deviations from Protontherapy Plan SETUP

Simulation Setup

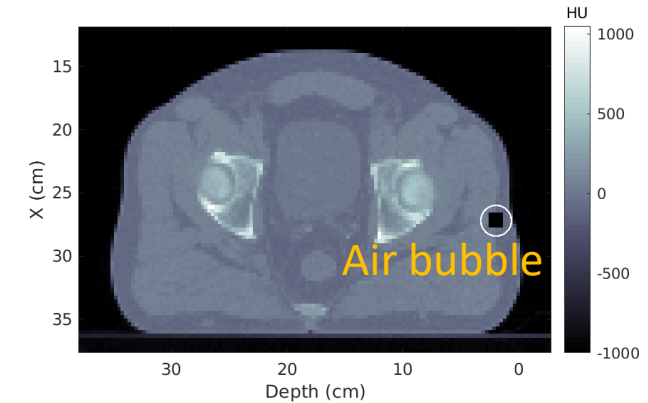


- Treatment plan for **prostate** tumor
- **4** ideal **detectors**
- Dictionary for axial **central slice**
- **GPU** k-Wave simulation (~ 90 min in NVIDIA GeForce RTX 2080 Ti)
- **Noise** added

Robustness Evaluation

Changes from the original plan:

1. Air bubble
2. Weight gain
3. 1 mm Shift in patient position
4. 3.5 % error in HU to relative stopping power conversion

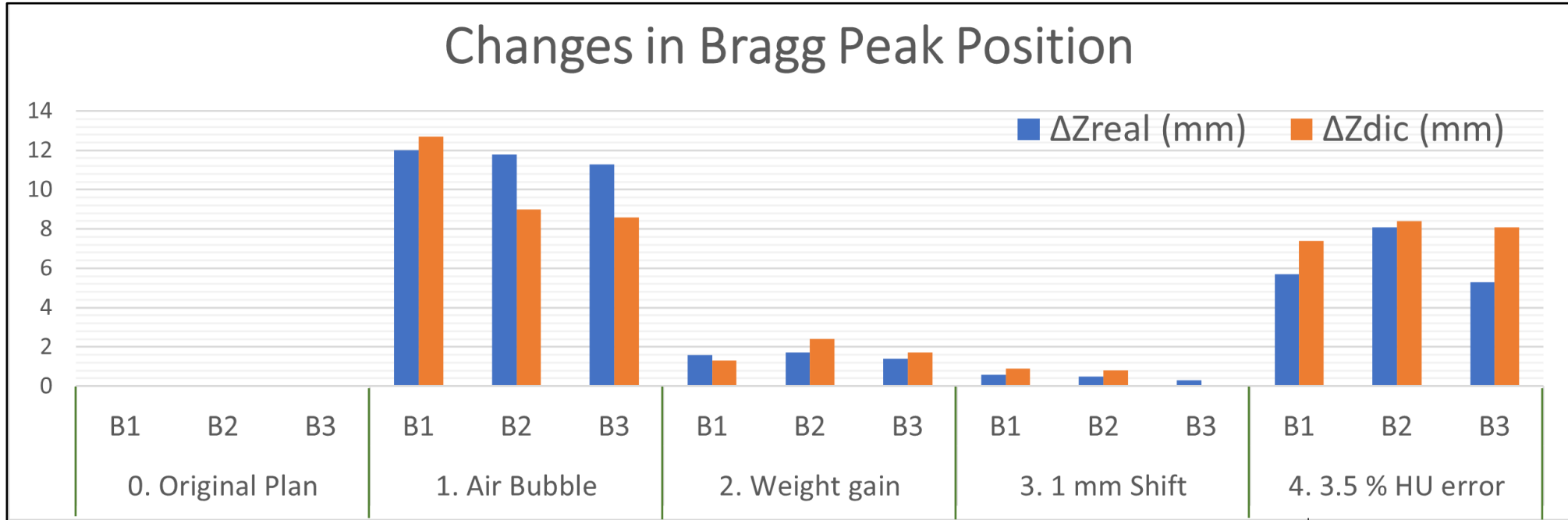


ΔZ_{real} Real change in BP position

VS

ΔZ_{dic} Change detected in BP position

2) Detection of Deviations from Protontherapy Plan RESULTS



Largest changes caused by air bubble and HU conversion error

The dictionary method detects the deviations in all cases

1.1 mm Average Accuracy (<0.7 %)

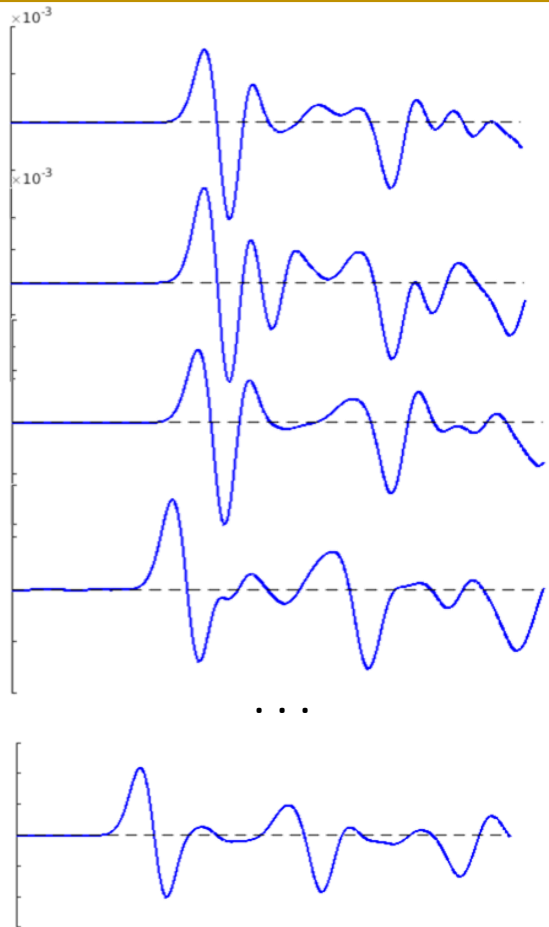
Dictionary-based protoacoustic dose map imaging for proton range verification. Clara Freijo et al. Photoacoustics, 21, 2021, 100240



It could alert of errors

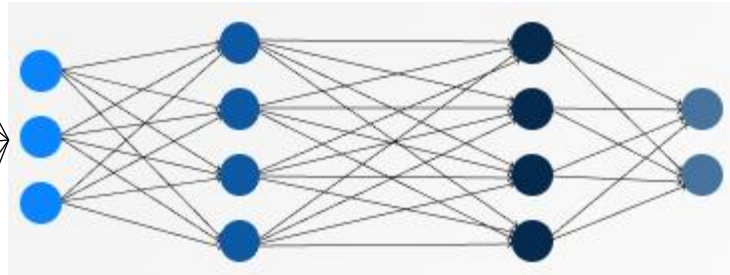
Method 3: Deep-Learning

Input: Acoustic signals



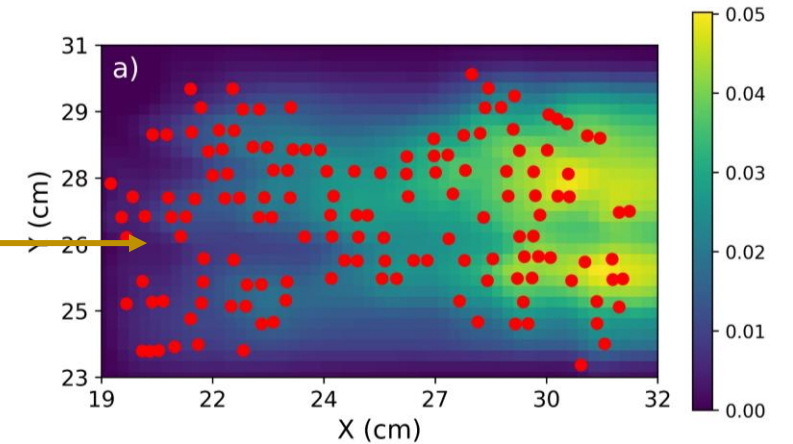
Neural Network

- 1.5M parameters
- 430 cases: 290 training / 140 validation



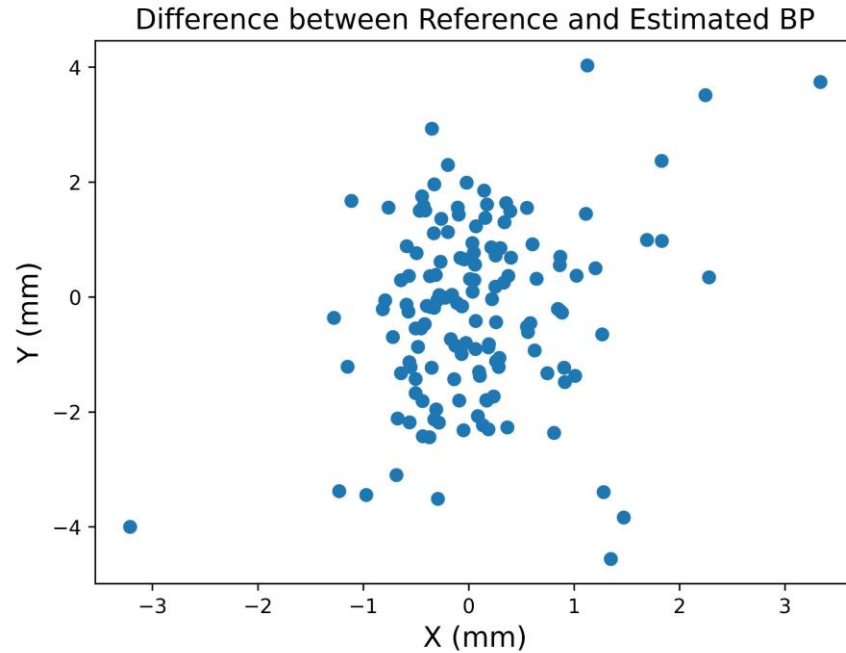
- Loss function: Mean Squared Error (MSE)
- Training time: 20 min

Output: Bragg Peak positions (X,Y)



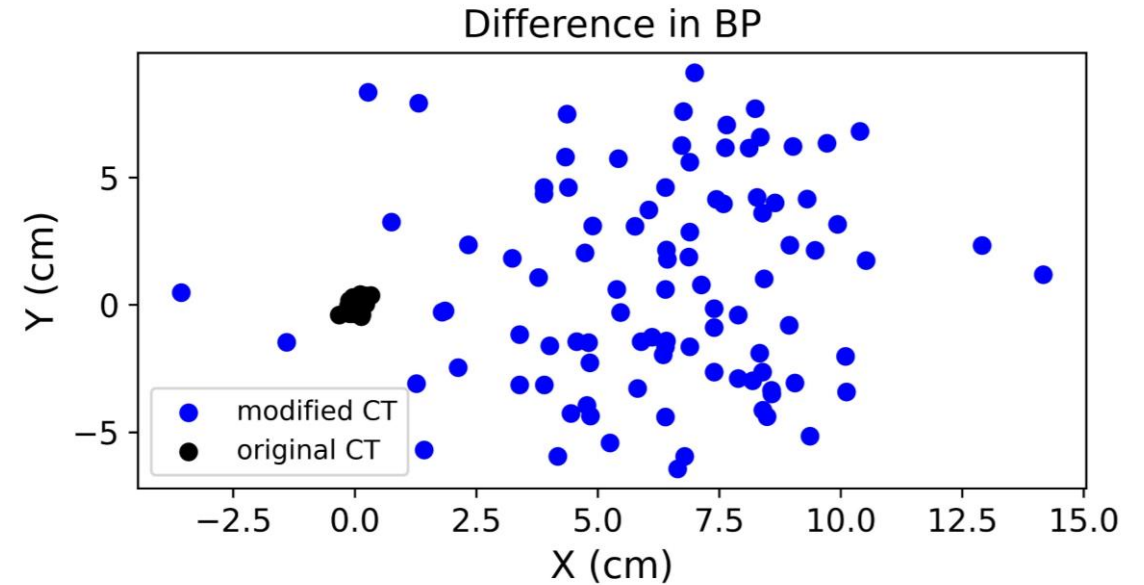
3) Protoacoustic Neural Network

Original CT



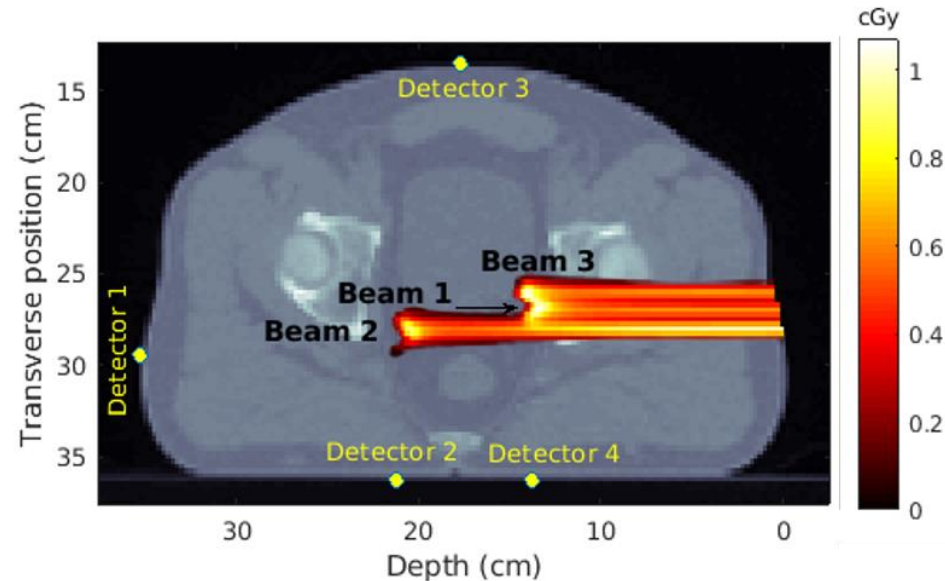
0.9 mm Average Accuracy

Modified CT: Air Region



Detection of deviations from the original BP position

- Real data will be acquired with phantoms at Hospital Quirón in Madrid in the following weeks.
- Combination of ultrasound imaging at low frequencies with protoacoustics may provide complementary information (anatomy and dosimetry).
- Combination of protoacoustic information with prompt-gamma and/or PET measurements.



- Novel approaches for proton range verification based on a **protoacoustic data**.
- A **proof-of-concept study** demonstrated the **feasibility** of implementation of the proposed method.
- Experiments in proton facilities to **test** the method in a **real scenario**.
- REFERENCES:

Iterative Reconstruction Methods in Protoacoustic Imaging" Book Chapter "Horizons in Computer Science Research V. 18" Nova Science Publishers may. 2020

Dictionary-based protoacoustic dose map imaging for proton range verification. Clara Freijo et al. Photoacoustics -Volume 21, March 2021, 100240



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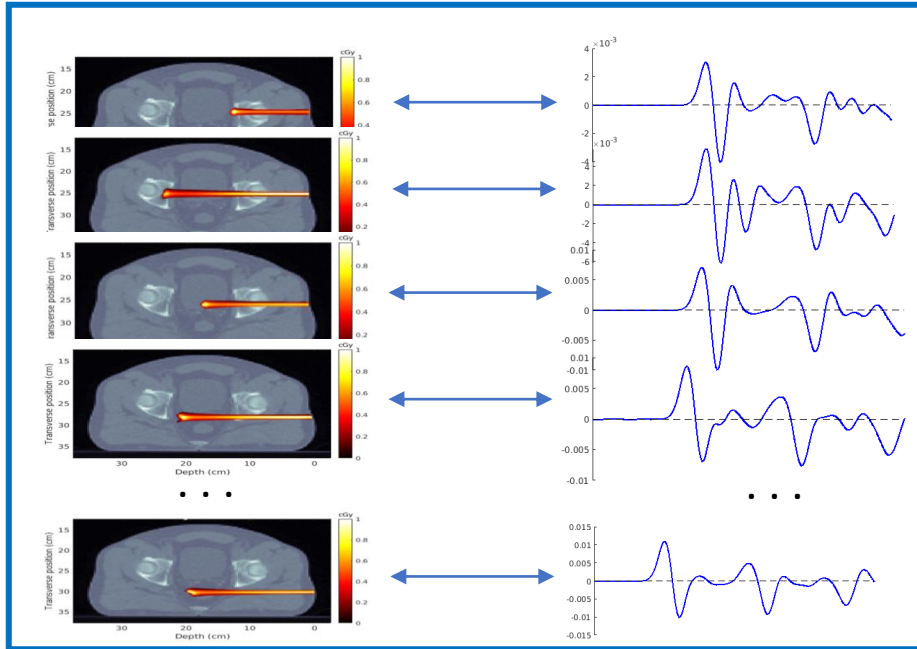


Plan de Recuperación,
Transformación y
Resiliencia



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