

# Optimizing the NEXI acquisition protocol for human gray matter microstructure mapping on a clinical MRI scanner using Explainable AI

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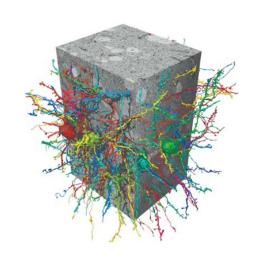
## Declaration of Financial Interests or Relationships

Speaker Name: Quentin Uhl

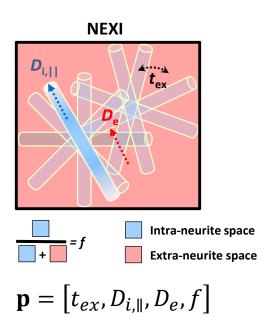
I have no financial interests or relationships to disclose with regard to the subject matter of this presentation.



#### The Neurite Exchange Imaging model



Grey matter microstructure (under the electron microscope)



- Geometry different from white matter
- Little myelin: water exchange across the cell membrane

#### From the parameters to the signal

| Kernel:                               | $\mathcal{K}(q, t, \mathbf{g} \cdot \mathbf{n}; f, D_{i,\parallel}, D_e, t_{ex}) = f' e^{-q^2 t D_i'} + (1 - f') e^{-q^2 t D_e'}$   |
|---------------------------------------|---|
| Where:<br>"apparent"<br>diffusivities | $D'_{i/e} = \frac{1}{2} \left\{ D_{i,\parallel} (\mathbf{g} \cdot \mathbf{n})^2 + D_e + \frac{1}{q^2 t_{ex}} \mp \left[ \left[ D_e - D_{i,\parallel} (\mathbf{g} \cdot \mathbf{n})^2 + \frac{2f - 1}{q^2 t_{ex}} \right]^2 + \frac{4f(1 - f)}{q^4 t_{ex}^2} \right]^{\frac{1}{2}} \right\}$ |
| "apparent"<br>fraction                | $f' = \frac{1}{D_i' - D_e'} \left[ f D_{i,\parallel} (\mathbf{g} \cdot \mathbf{n})^2 + (1 - f) D_e - D_e' \right]$  |
| Powder average (over directions):     | $\bar{S}(\mathbf{p},q,t) = S \Big _{q=0} \cdot \int_0^1 \mathcal{K}(q,t,\mathbf{g} \cdot \mathbf{n};\mathbf{p}) d(\mathbf{g} \cdot \mathbf{n})$   |

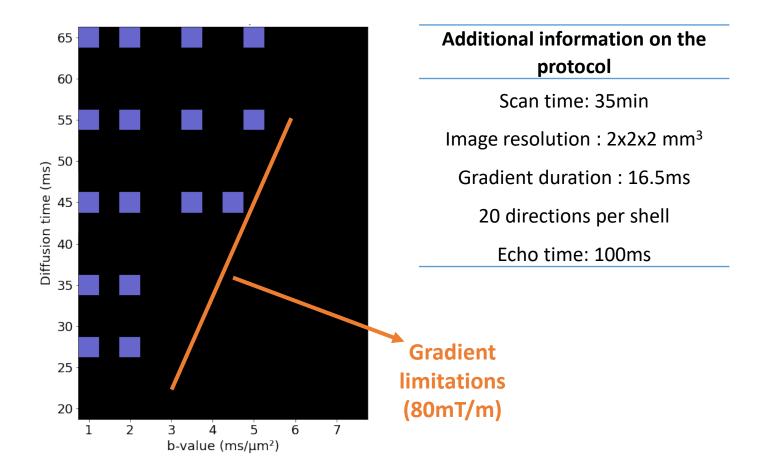
$$b = q^2 t = (\gamma G \delta)^2 t$$



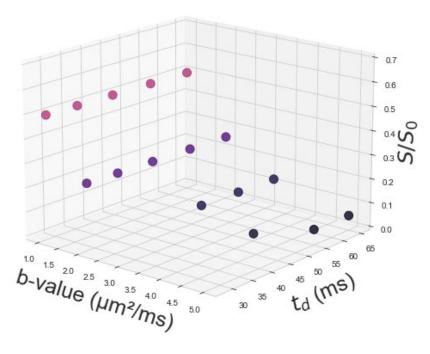
multiple acquisition parameter combinations



#### Our MRI Protocol on the clinical 3T Prisma Scanner

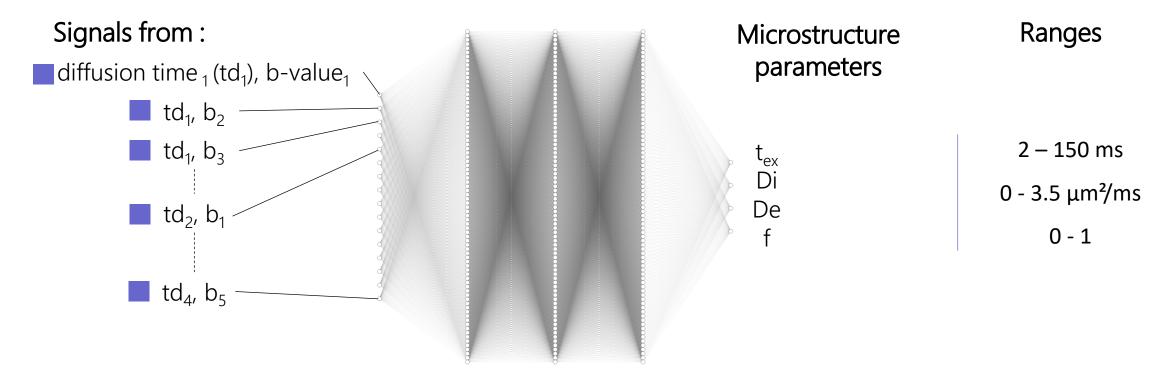


## Powder-averaged signal S depends on b and t<sub>d</sub>





#### Retrieving parameters using a Multi-Layer Perceptron (MLP)



3 hidden layers, 500 neurons each



#### Retrieving parameters using a Multi-Layer Perceptron (MLP)

Training set: 2 000 000 examples

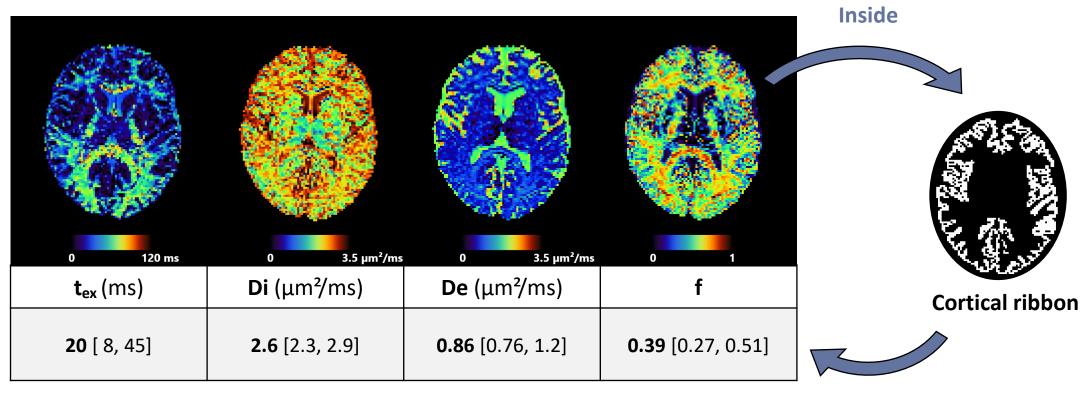
Test set: 10 000 examples

SNR: 80-120

|   | t <sub>ex</sub> (ms) | Di (μm²/ms) | <b>De</b> (μm²/ms) | f     |
|---|----------------------|-------------|--------------------|-------|
| Clinical Protocol Mean Absolute Error (MAE) | 27.457               | 0.423       | 0.351              | 0.083 |



#### MLP estimates on an axial slice of the human brain

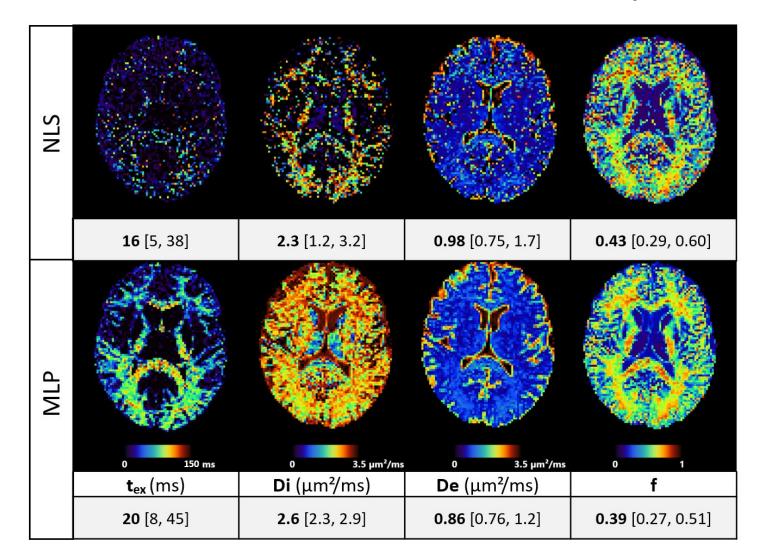


Statistics on the cortical ribbon

Median [1st, 3rd quartile] of cortical map

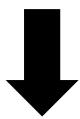


#### Comparison with the Non-linear Least Squares estimates





How to improve this protocol to get lower errors?



Explainable AI

**Shapley value** of input i ~ its contribution to the output



#### Explainable AI using Shapley values

Shapley value of player i ~ his/her contribution to the game:

$$\varphi_i = \frac{1}{\# \ of \ players} \sum_{coalitions \ excluding \ i} \frac{marginal \ contribution \ of \ i \ to \ coalition}{\# \ of \ coalitions \ excluding \ i} \ \# \ of \ coalitions \ excluding \ i \ of \ this \ size$$

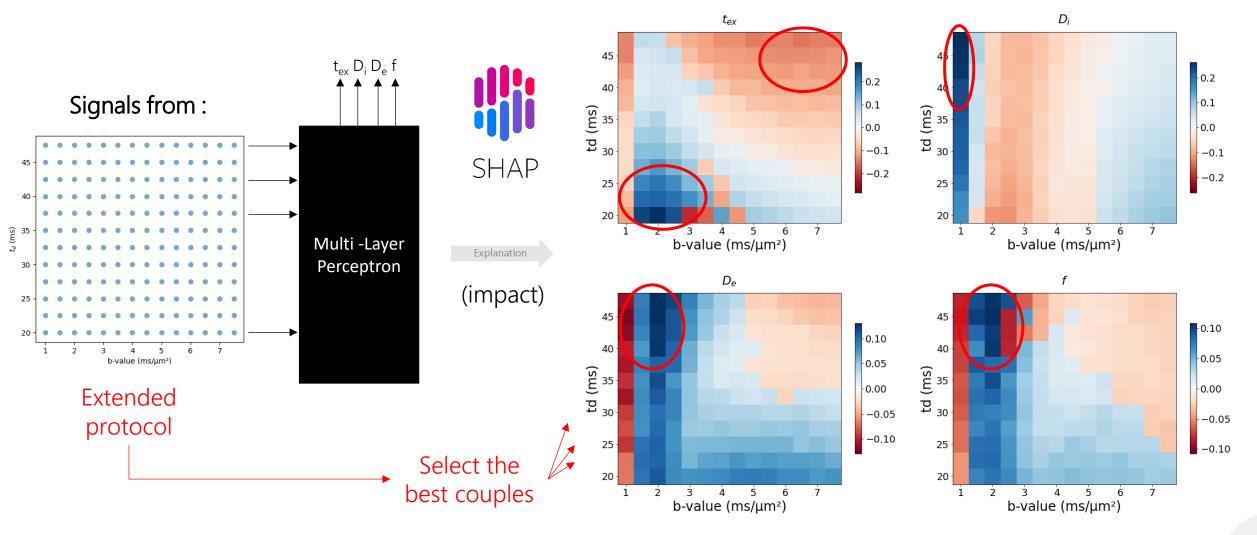
the score of the team with the player

the score of the team without the player



#### Looking for an optimized protocol

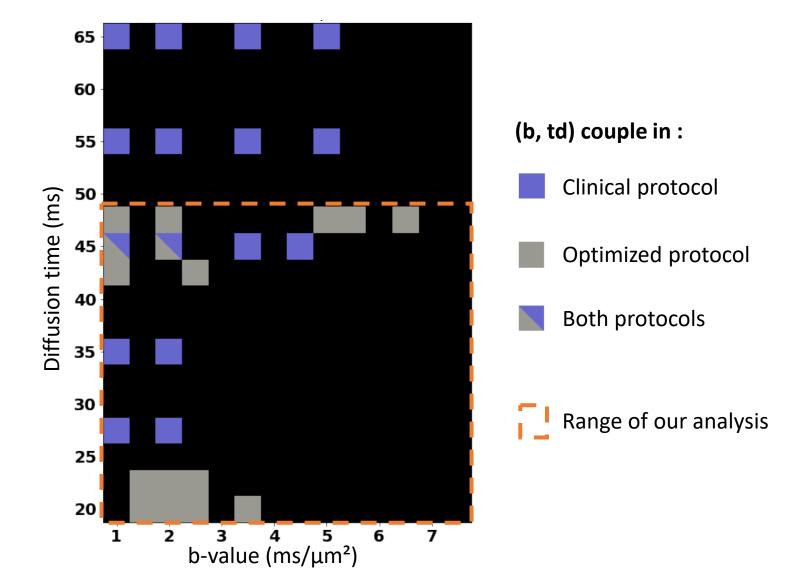
Sign(Median(SHAP value)) x Mean(|SHAP value|) per (b, td) couple for each parameter



Source: SHAP, Scott Lundberg



#### The two protocols





## Is this new protocol really better?

|                           | t <sub>ex</sub> (ms) | <b>Di</b> (μm²/ms) | De (μm²/ms) | f      |
|---------------------------|----------------------|--------------------|-------------|--------|
| Clinical Protocol MAE     | 27.457               | 0.423              | 0.351       | 0.083  |
| Optimized Protocol<br>MAE | 25.496               | 0.419              | 0.330       | 0.076  |
|                           | -1.961               | -0.004             | -0.021      | -0.007 |

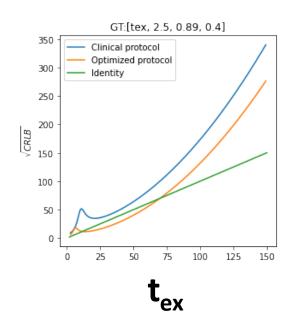
→ **A** Still dependent on the training & test sets

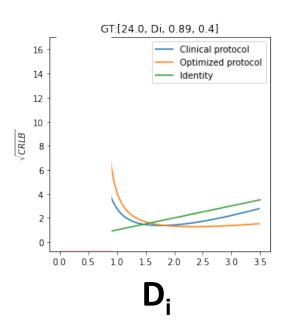


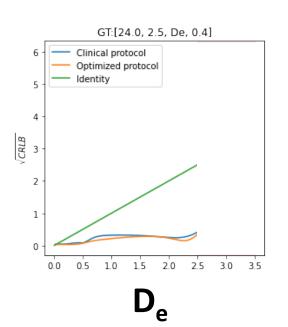
## Is this new protocol really better?

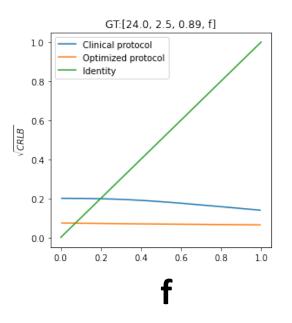
To compare protocols, we set side by side their associated **Cramer-Rao Lower Bound (CRLB)** on some sets of microstructure parameters.

#### Around [24, 2.5, 0.89, 0.4]









▲ Depends on the parameters p (4D)



#### Summary & Perspectives

- ➤ We succeeded in estimating NEXI parameters on the human brain on a clinical scanner.
- > A naïve protocol seems to already hold reasonable results.
- ➤ Next, we will identify a new protocol with this method within a range adapted to the Prisma gradient limitations, implement it experimentally and compare the maps.



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