

# PINN for solving PDE and Its application in Alzheimer's disease

## Abstract

- Alzheimer's disease (AD) is a neurodegenerative disorder characterized by the accumulation of misfolded tau proteins in the brain, crucial to disease progression. This study integrates Physics-Informed Neural Networks (PINNs) with symbolic regression to model the dynamics of tau proteins using reaction-diffusion equations. By applying our approach, we successfully predicted the reaction term in the equation, revealing a faster rate of tau misfolding in individuals with Alzheimer's compared to healthy controls.

## Problem definition

- Alzheimer's disease (AD) involves the accumulation of misfolded tau proteins in the brain, which can be modeled using a reaction-diffusion equation (PDE). The concentration of tau proteins  $c(t,x)$  evolves over time, described by:

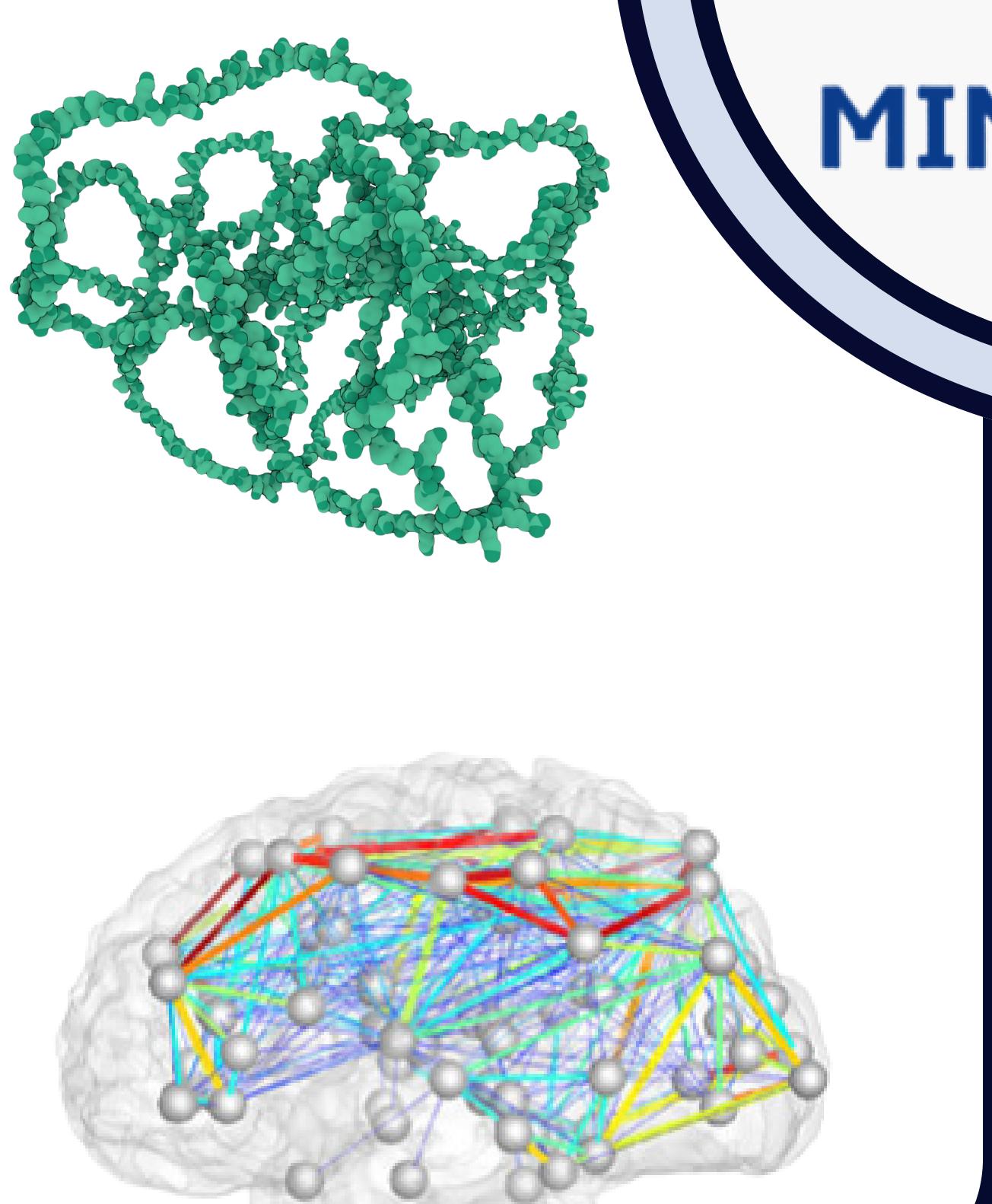
$$\frac{\partial C}{\partial t} = \nabla \cdot (D \cdot \nabla C) + f(C)$$

- We discretize the reaction-diffusion model on a weighted graph  $G$ , where each node  $i$  represents a brain region and the edges represent connections between them. This gives a discretized equation for the change in tau protein concentration  $c_i$  at each region over time:

$$\frac{dc_i}{dt} = h_i^k(t, C) + f(c_i) \rightarrow i = 1, \dots, N$$

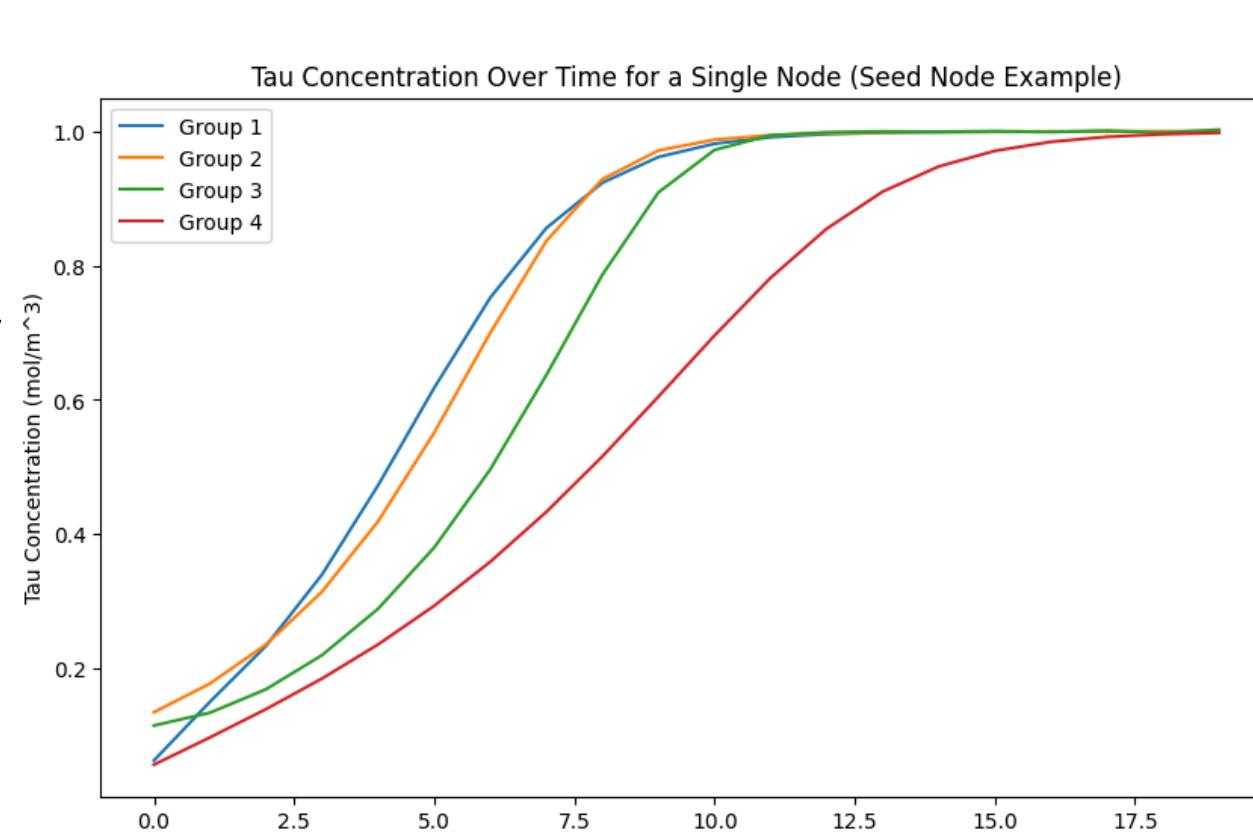
## Literature review

- Chronological Structure: Early studies focused on predefined reaction terms to model biological processes. But in our approach we use multiple reaction terms for more precise prediction.
- Methodological Structure: (PINNs) Proven effective for solving PDEs with missing terms, though applications to tau protein dynamics are still emerging.



## Data description

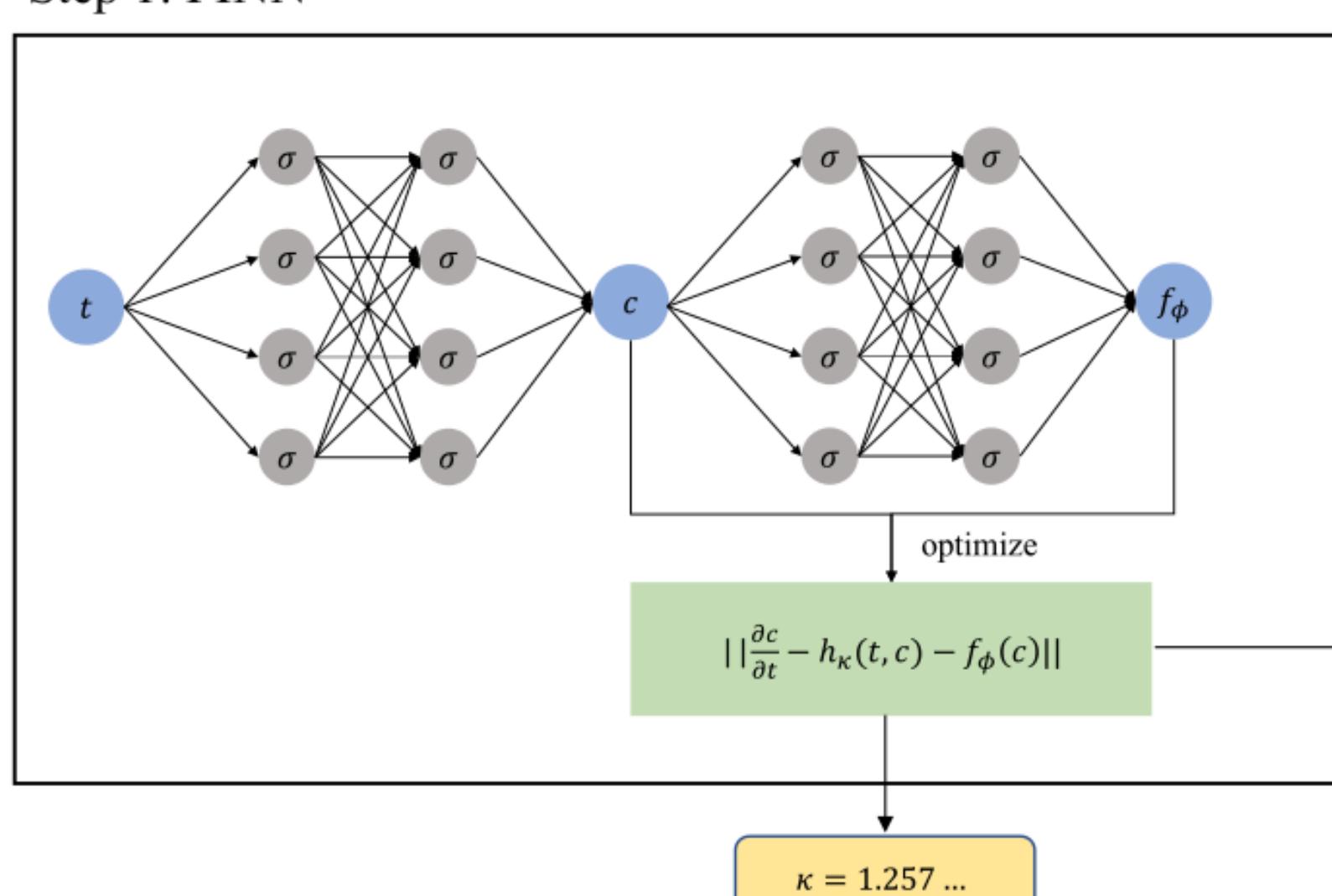
- In this study, we modeled tau concentration for 76 subjects, divided into four groups, using synthetic data.
- Parameters  $k$ ,  $\alpha_i$ , and  $\alpha_{ij}$  were assigned distributions  $\text{BoundNormal}(1, 0.52)$ ,  $N(0.6, 0.12)$ , and  $N(\alpha_i, 0.22)$ , respectively.
- Initial tau concentrations,  $c_i(0)$ , were sampled from a normal distribution based on real data.



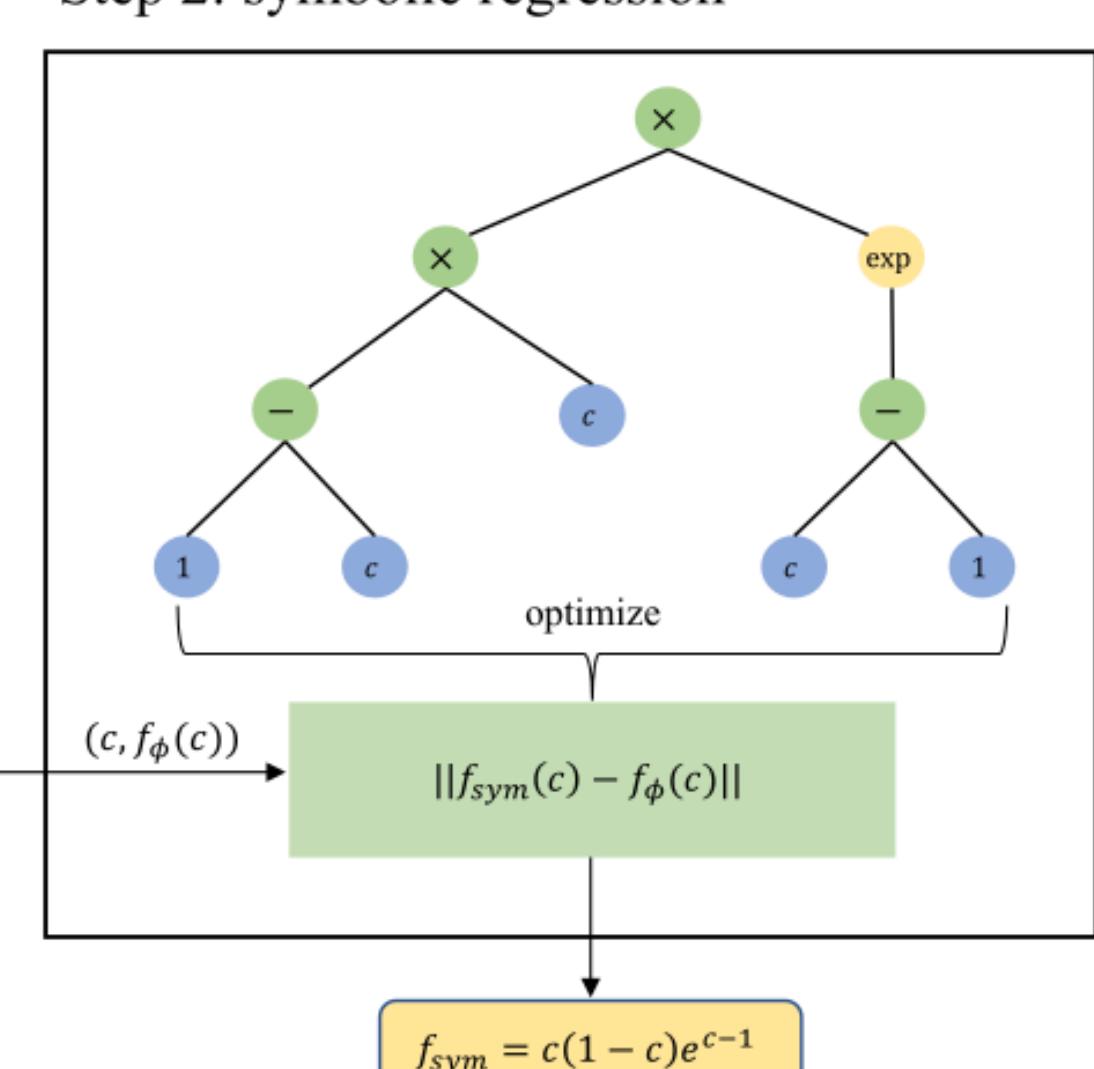
## Methodology

- Our approach consists of two main branches PINN and Symbolic Regression.

Step 1: PINN



Step 2: symbolic regression



- For discovering  $f$  is formulated as follows: first infer  $c$ ,  $f$ , and  $\kappa$  from data of  $c$  and the physics defined in this equation using PINNs, and then find the analytic expression of  $f$  using symbolic regression.

$$\frac{dC_i}{dt} = -K \sum_{j=1}^N L_{ij} C_j + \alpha f(C_i) \rightarrow i = 1, \dots, N$$

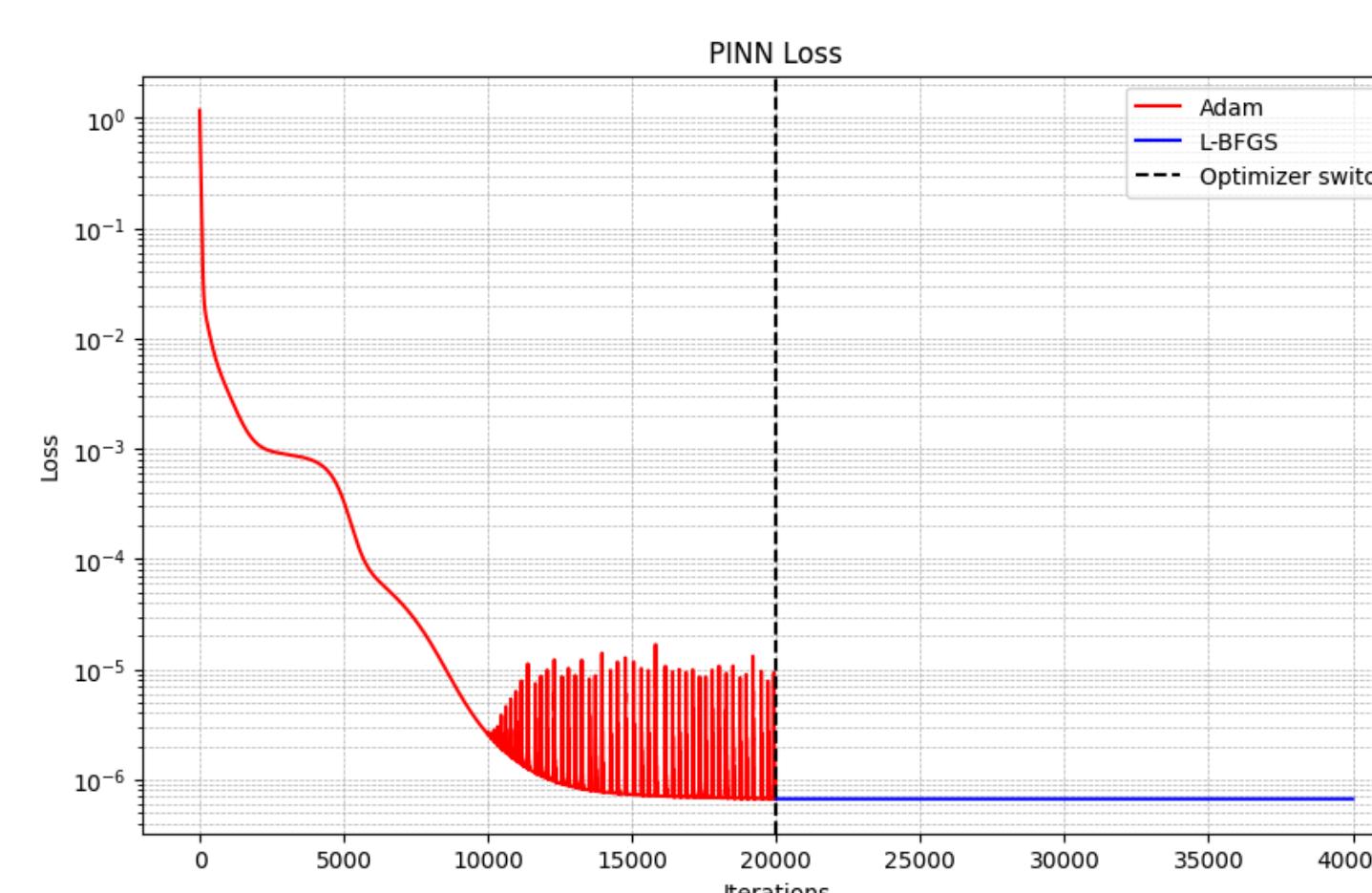
## Results



- PINNs and symbolic regression accurately predicted tau protein concentrations, demonstrating alignment with synthetic data.

- Integrated physical laws into the loss function for efficient training without numerical methods.
- Used Adam Optimizer (20,000 epochs) for initial training and L-BFGS Optimizer (20,000 epochs) to ensure convergence.

- Identified interpretable reaction term  $f(c)$  using symbolic regression.



$\kappa$	$\alpha$	$f_{\text{sym}}$
0.29	0.15	$8.908 \cdot c^3 - 23.446 \cdot c^2 + 15.667 \cdot c + 0.792$

## Conclusion

- We demonstrated the effectiveness of combining PDE-based modeling with machine learning.
- We predicted tau concentrations and patterns using synthetic data.
- We used symbolic regression and PINNs to model tau dynamics in Alzheimer's.

## Future work

- Enhance tau dynamics model with PET and MRI data.
- Visualize tau trends and disease progression using a web-based platform.

