



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:

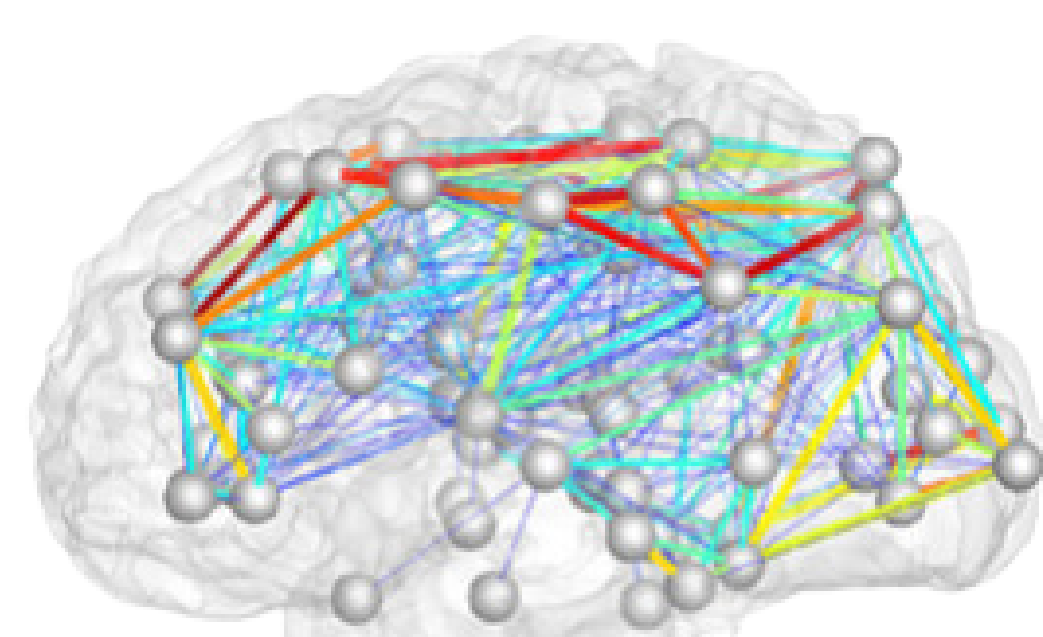
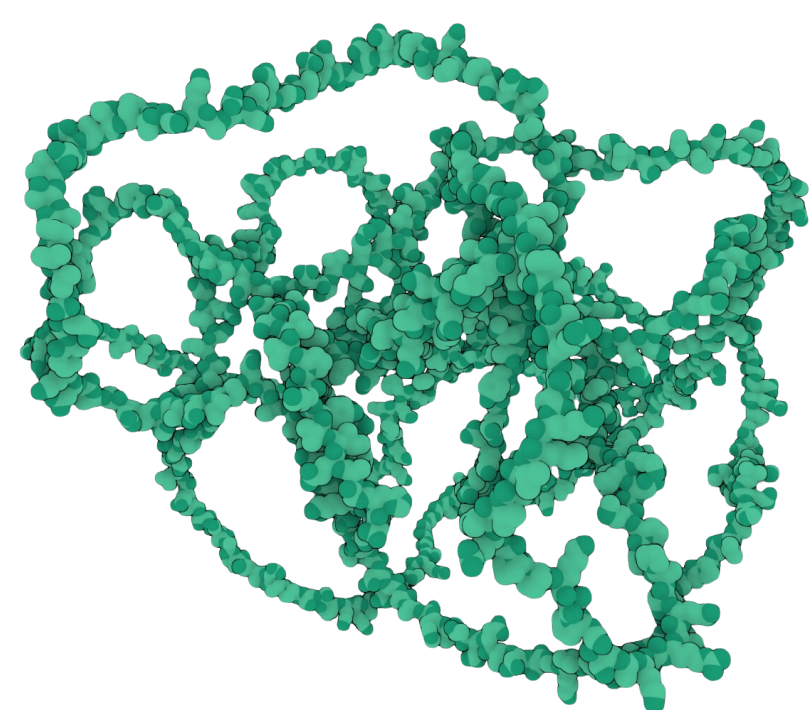
$$\frac{\partial C}{\partial t} = \nabla \cdot (D \cdot \nabla C) + f(C)$$

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

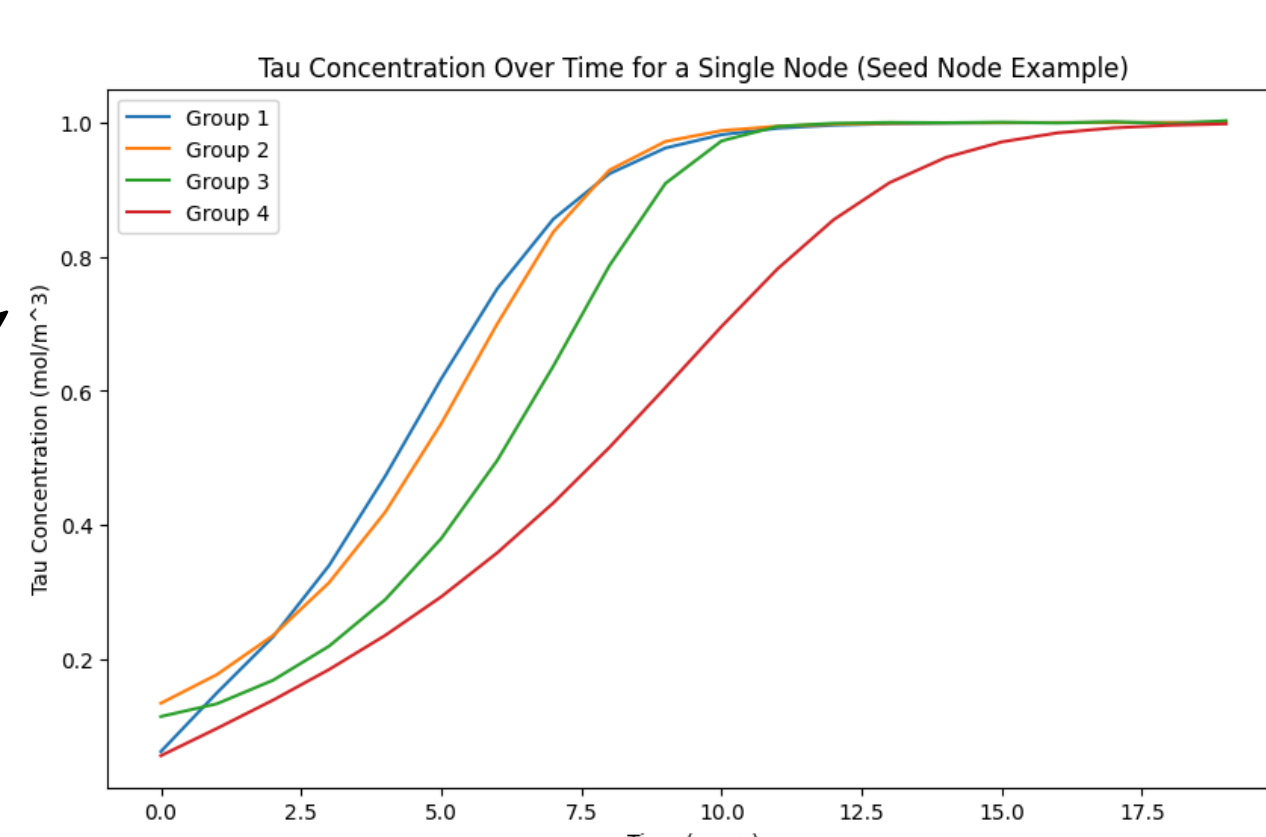
- Chronological Structure:** Early studies focused on predefined reaction terms to model biological processes, But in our approach we use multiple reaction terms for more precise prediction.
- Methodological Structure:** (PINNs) Proven effective for solving PDEs with missing terms, though applications 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 , α_i , and α_{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.

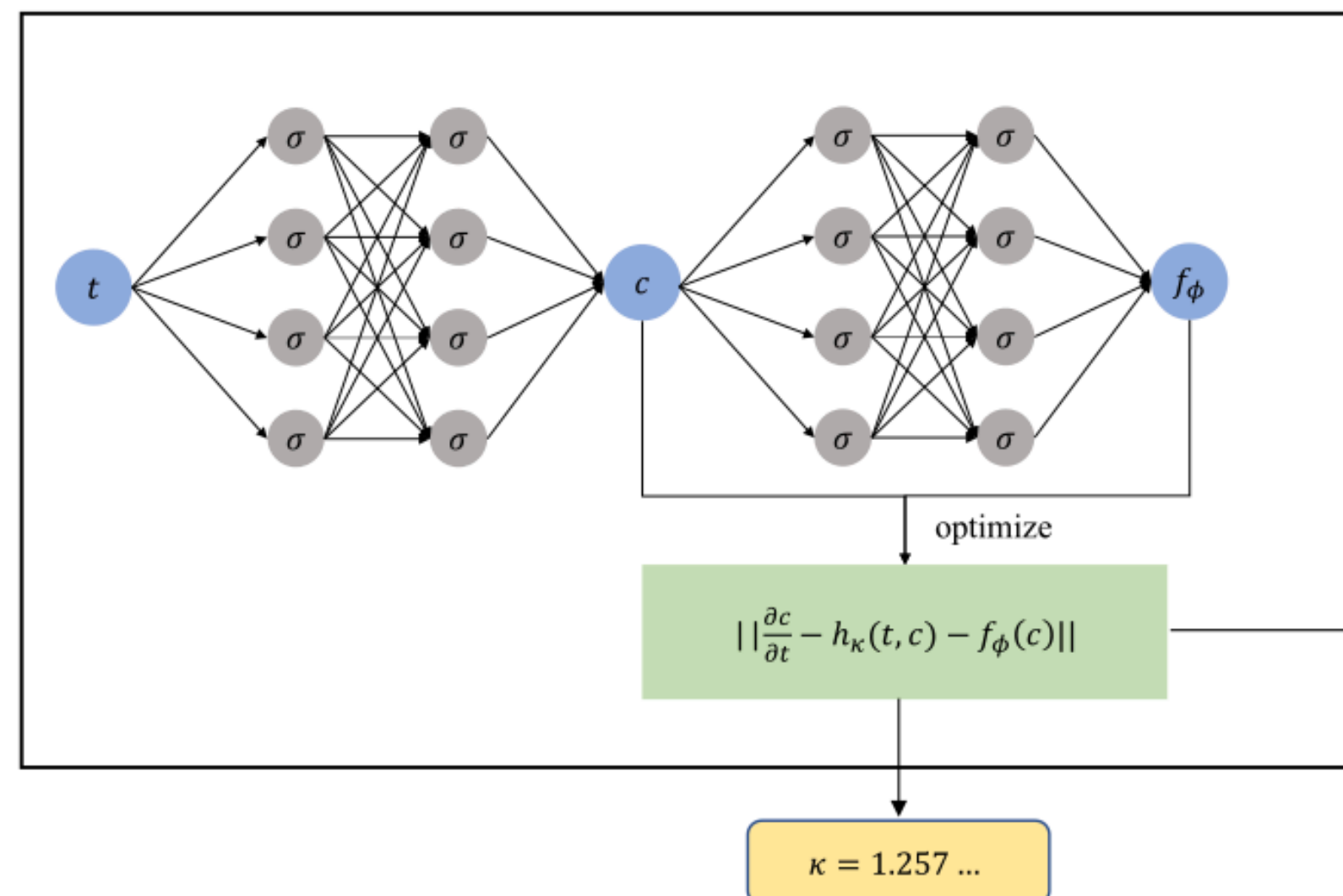
- Tau concentrations were sampled at $t_k = k$ for $k = 0$ to 20 years.



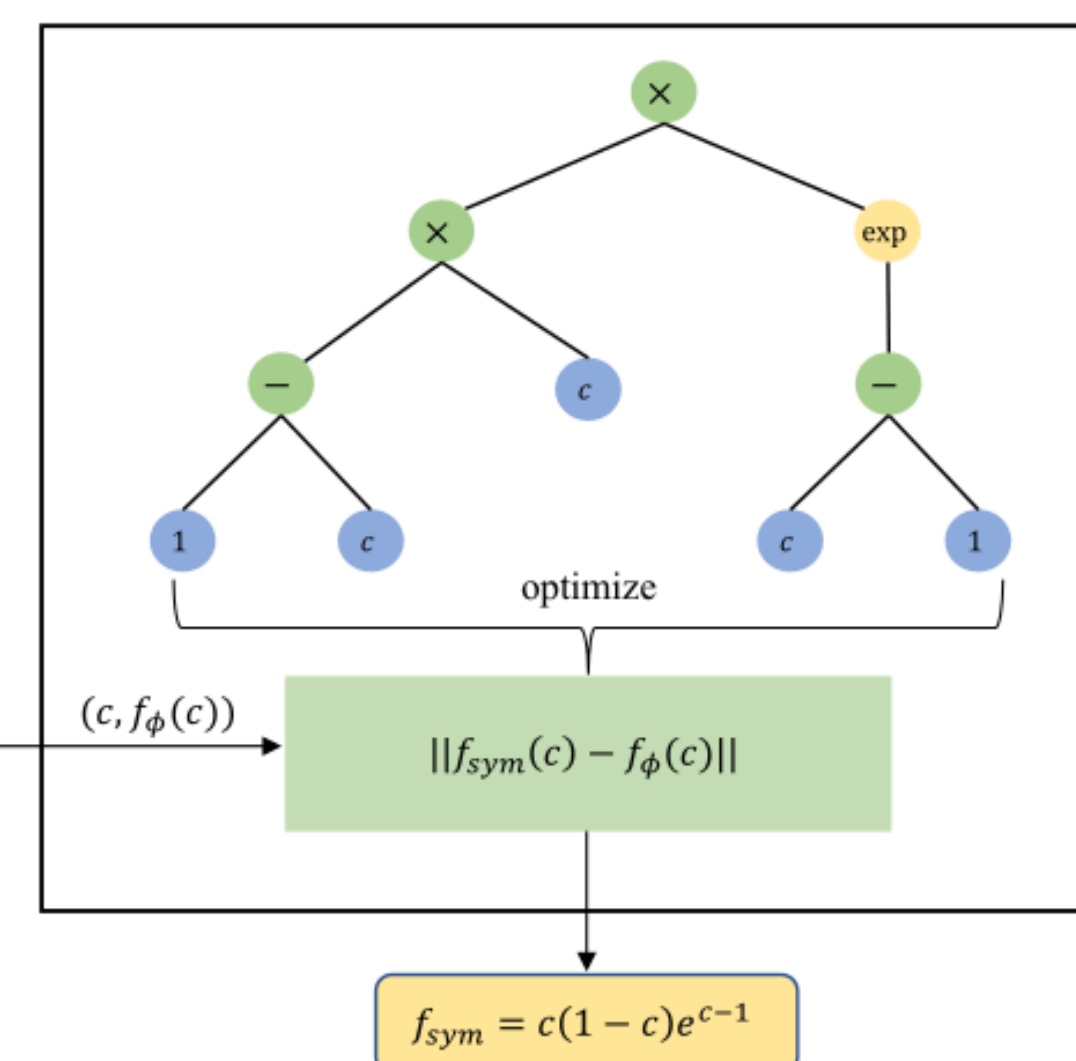
Methodology

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

Step 1: PINN



Step 2: symbolic regression

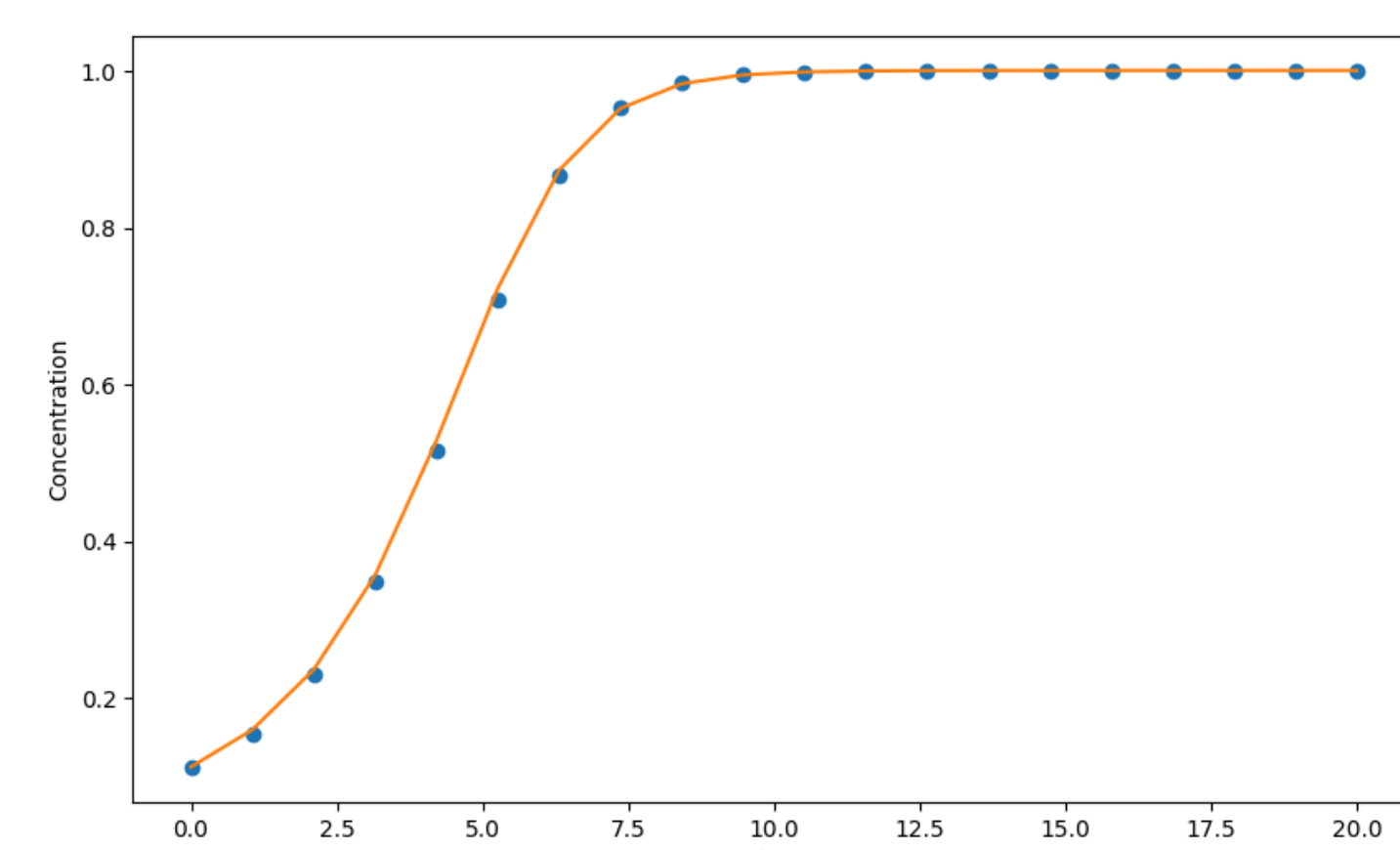
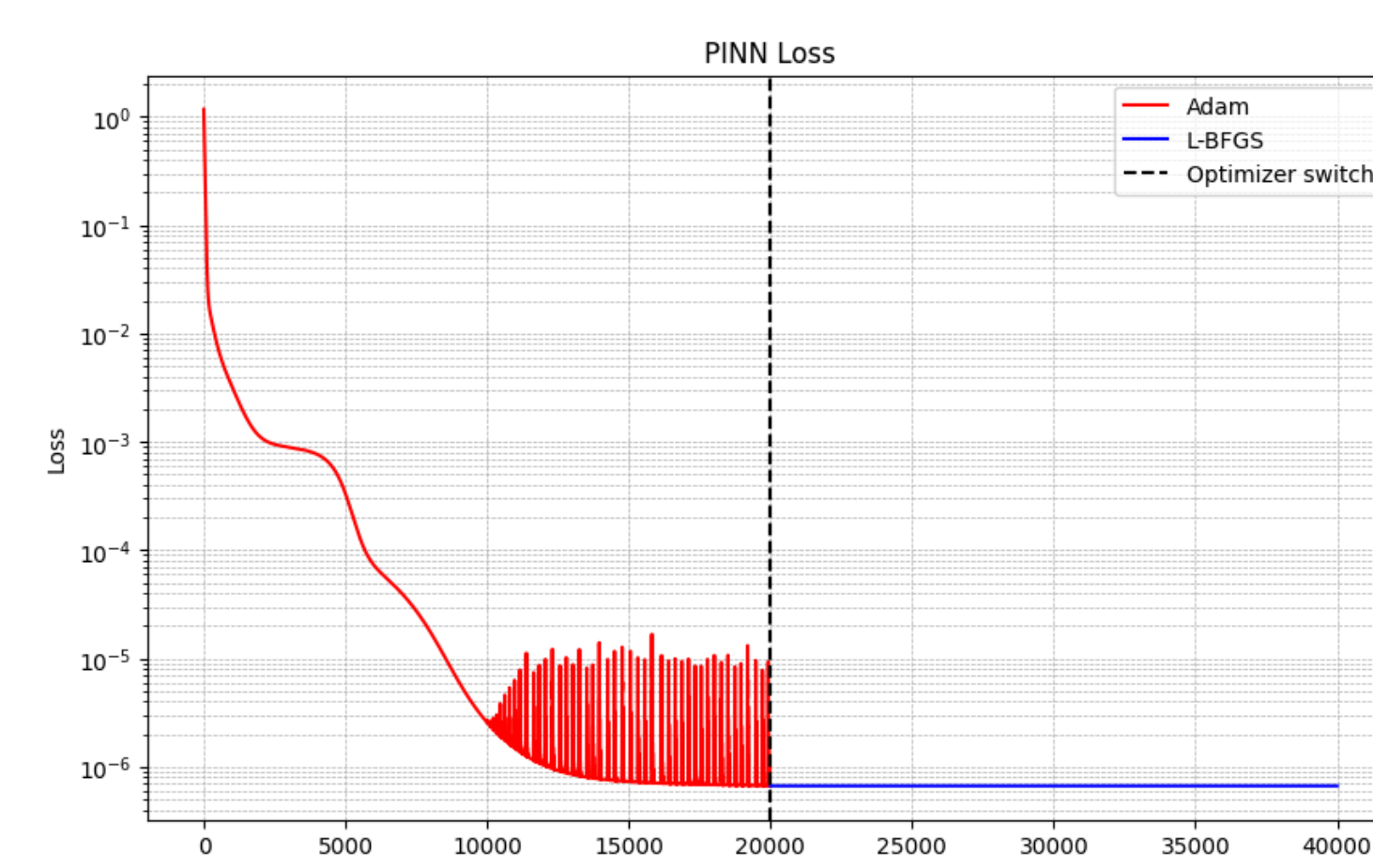


- For discovering f is formulated as follows: first infer c , f , and κ from data of c and the physics defined in this equation using PINNs, and then find the analytic expression of f using symbolic regression.

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

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.



κ	α	f_{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.

