

Action potential propagation in long-range axonal fibre bundles

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Introduction

Background

- Recent advances in MRI imaging make it possible to determine white matter structure non-invasively
- Action potential velocity determines delays between brain regions
- Standard numerical techniques to model the dynamics of action potentials require huge computational effort

Aims

- Develop a computationally efficient framework to compute action potential velocity (spike-diffuse-spike model)
- Determine how action potential velocity depends on structural parameters
- Study interaction of action potentials between ephaptically coupled fibres

Methods

- Axonal membrane potential V is described by cable equation

$$\tau \frac{\partial V}{\partial t} = \lambda^2 \frac{\partial^2 V}{\partial x^2} - V + R_m I_{chan}(V, t)$$

- We model channel currents I_{chan} as threshold-activated events (Fig. 1)

- The resulting linear cable equation can be solved explicitly

$$V(x, t) = \sum_{n=1}^N \Phi_I(x + nL, t + nt_{sp}) \quad \Phi_I(x, t) = R_m \int_0^t I_{chan}(t-s)G(x, s)ds$$

- Unknown time delay t_{sp} is computed numerically from

$$V_{thr} = \sum_{n=1}^N \Phi_I(nL, nt_{sp})$$

- Velocity is given by $v = L / t_{sp}$ (Fig. 2)

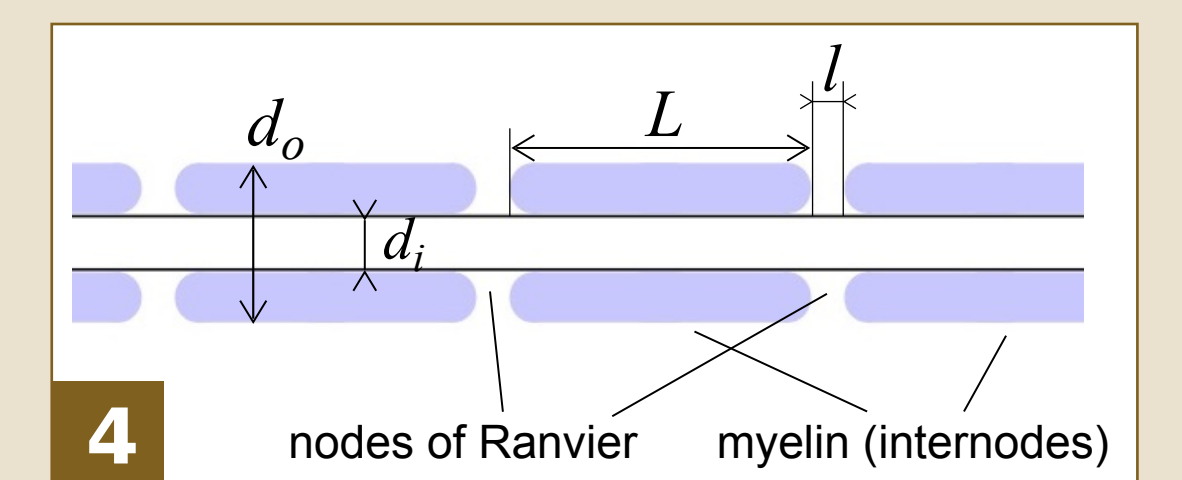
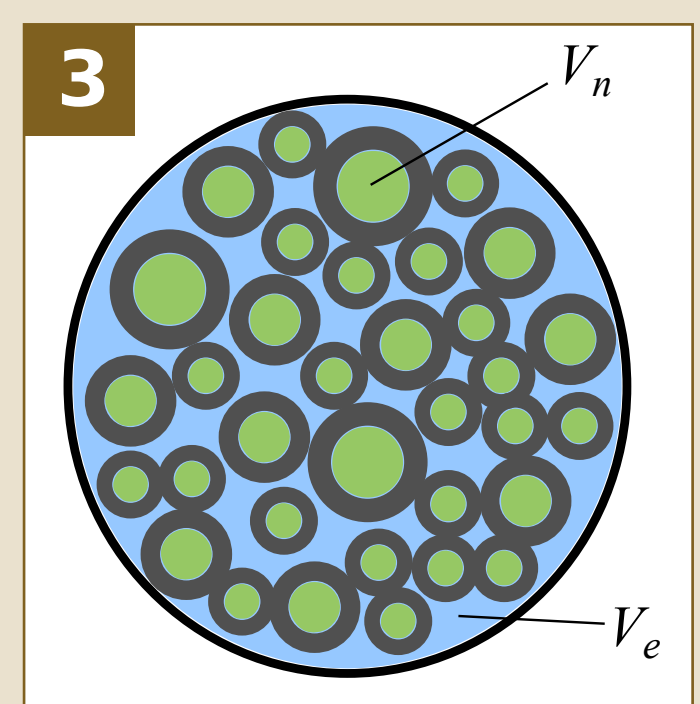
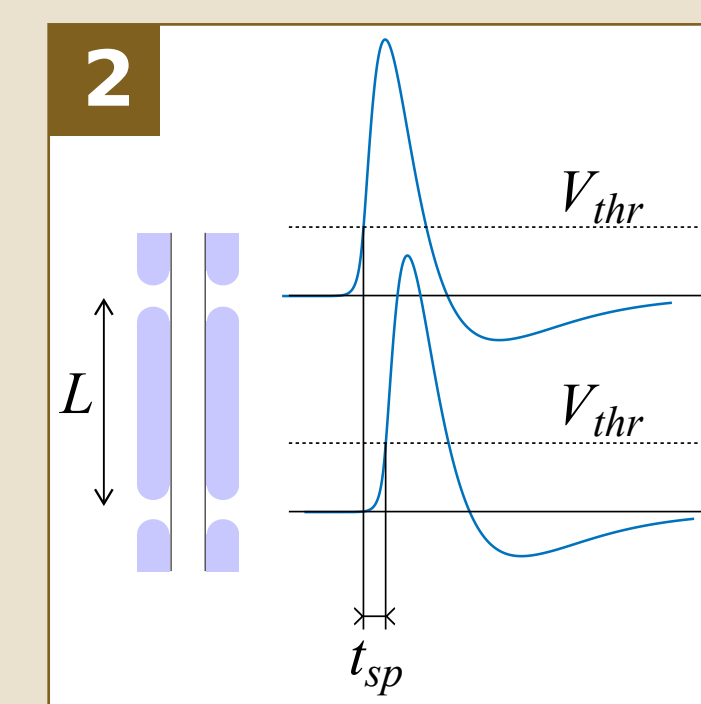
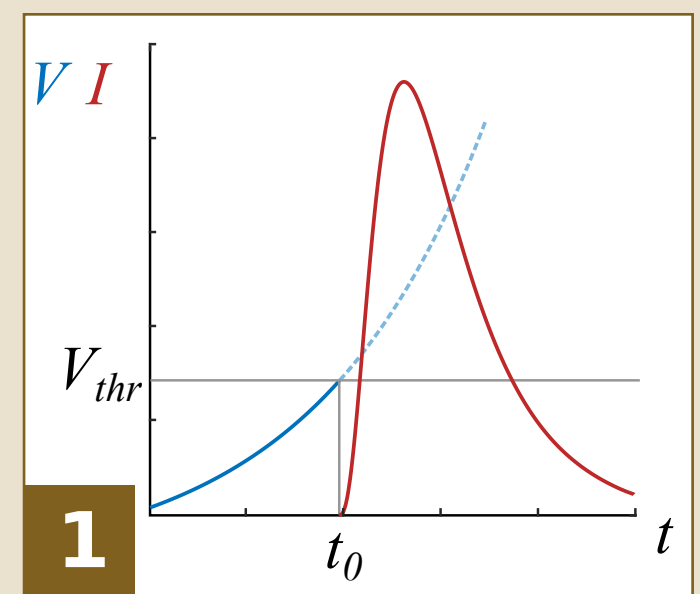
- Results are compared with detailed numerical model [1]

- Methods can be extended to axonal fibre bundles (Fig 3):

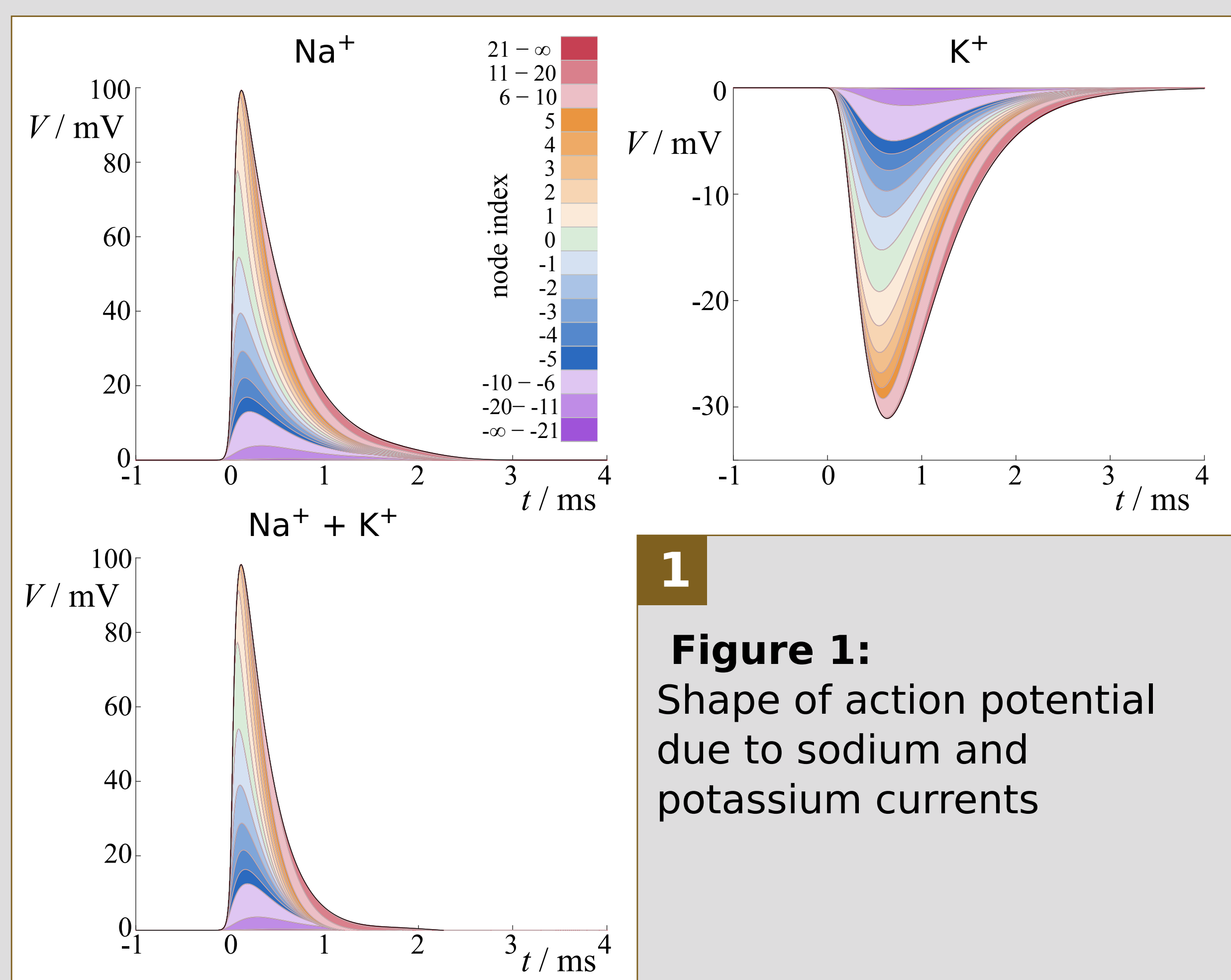
$$\tau \frac{\partial(V_n - V_e)}{\partial t} = \lambda_n^2 \frac{\partial^2 V_n}{\partial x^2} - (V_n - V_e) + R_m I_n^{chan}(t)$$

- Using assumptions from [2] and $U_n = V_n - V_e$

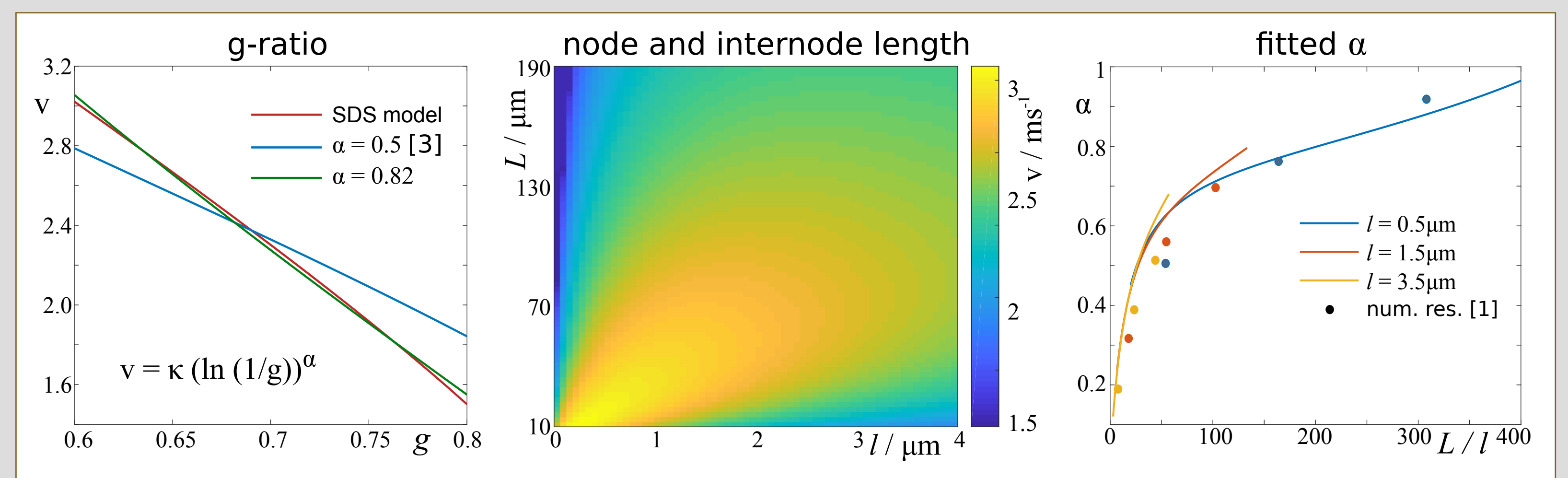
$$\tau \frac{\partial U_n}{\partial t} = \lambda_n^2 \frac{\partial^2 U_n}{\partial x^2} - U_n + R_m I_n^{chan}(t) - \alpha \lambda_n^2 \sum_p R_{ax,p}^{-1} \frac{\partial^2 U_p}{\partial x^2}$$



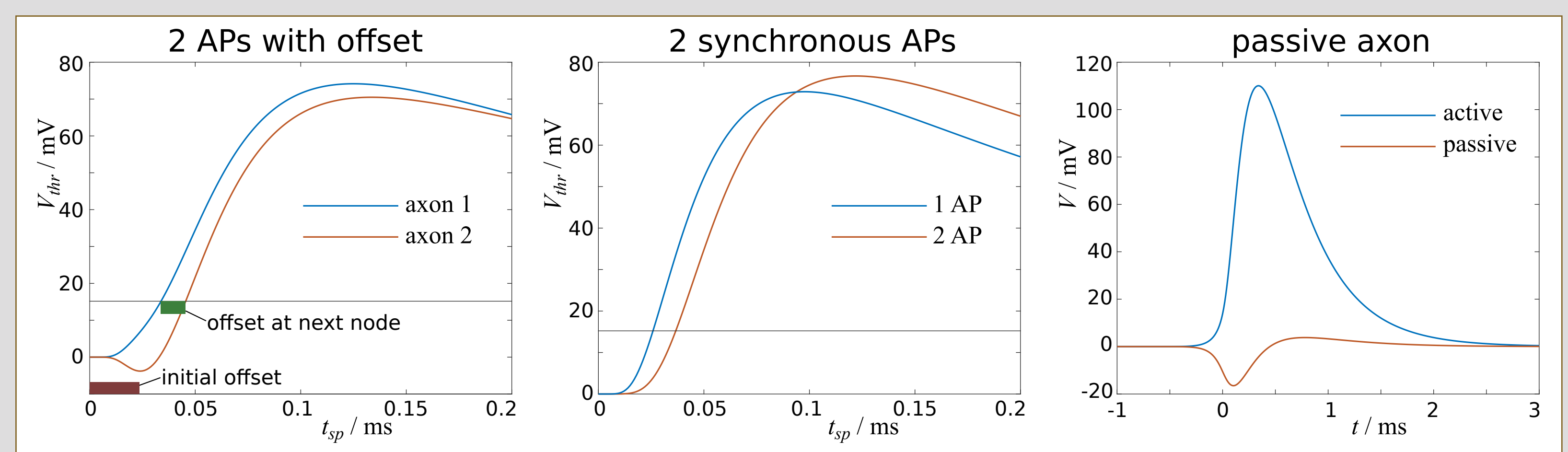
Results



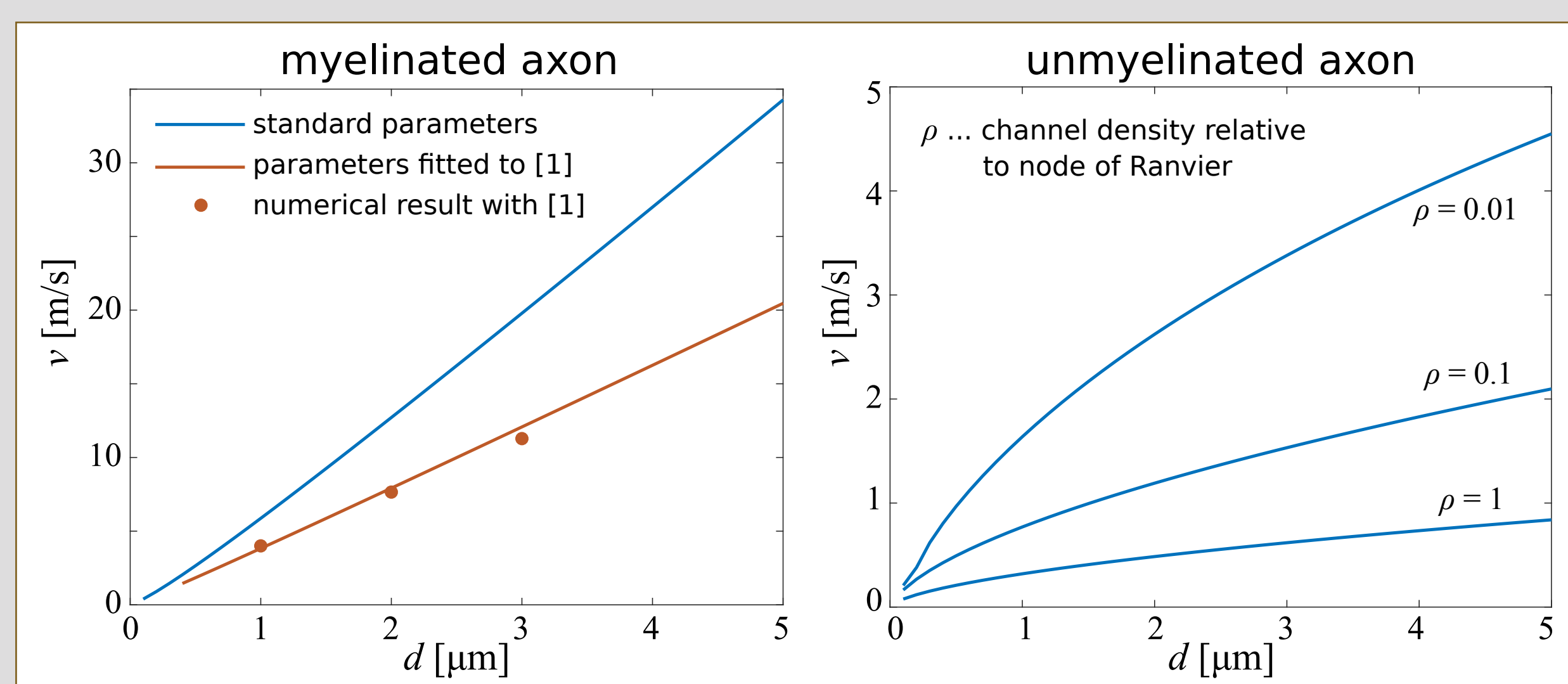
1 Figure 1: Shape of action potential due to sodium and potassium currents



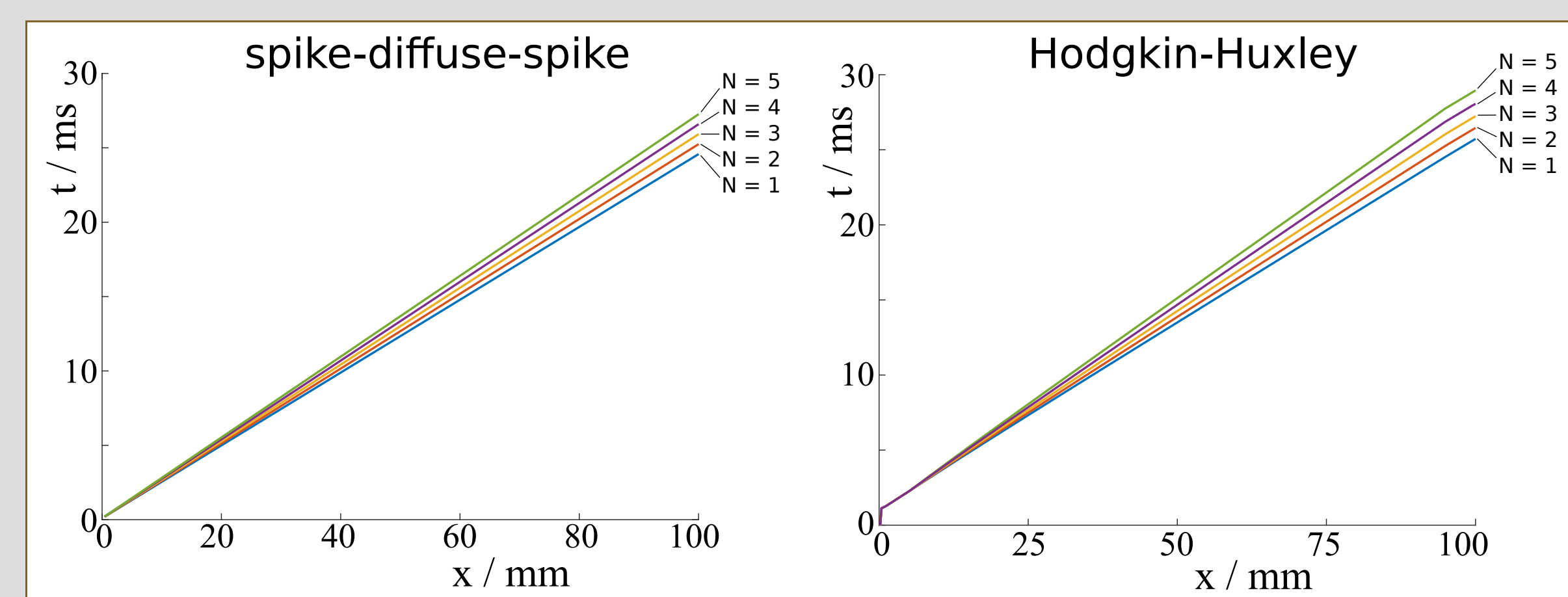
3 Figure 3: Propagation velocity and myelin structure



4 Figure 4: Action potential dynamics in a coupled pair of identical axons



2 Figure 2: Propagation velocity and axon diameter



5 Figure 5: Trajectories of spike volleys (N spikes) in fibre bundle with finite extra-axonal space

Conclusion

single axon

- We use a simplified description of action potential propagation without explicit Hodgkin-Huxley dynamics
- Approach yields an analytical expression for the (implicit) relationship between action potential velocity and model parameters
- Many known results are reproduced qualitatively and quantitatively
- velocity depends more strongly on g-ratio than inferred in [3]
- Speed-up by three orders of magnitude in comparison to numerical integration of cable equation

axon bundle

- Ephaptic coupling due to limited extra-axonal space synchronises and slows down action potentials (as observed by [2] and [4])
- Ephaptic coupling leads to activity-dependent delays

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