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## Models of Neural Systems, WS 2019/20

### Project 2: Action potential propagation

Project presentation and report submission: February, 11th, 2020

#### Background

The aim of this project is to model the propagation of an action potential along an axon. The relationship between the membrane current  $i_m$  and the voltage  $V$  along an axon is given by the equation

$$c_m \frac{\partial V}{\partial t} = \frac{1}{2ar_L} \frac{\partial}{\partial x} \left( a^2 \frac{\partial V}{\partial x} \right) - i_m + i_e, \quad (1)$$

where  $a$  is the radius of the axon,  $r_L$  is the intracellular resistivity and  $c_m$  the specific membrane capacitance.

The ionic current  $i_m$  flowing through a patch of axonal membrane is well-described by the Hodgkin-Huxley model:

$$i_m = \bar{g}_L(V - E_L) + \bar{g}_{Na}m^3h(V - E_{Na}) + \bar{g}_Kn^4(V - E_K), \quad (2)$$

where  $m$ ,  $h$  and  $n$  are Hodgkin-Huxley-type gating variables.

Combining the two equations leads to a partial differential equation that can be solved numerically by a multi-compartmental approximation. In a nonbranching cable, each compartment  $\mu$  is coupled to two neighbors, and the equations for the membrane potentials of the compartments are

$$c_m \frac{dV_\mu}{dt} = -i_m^\mu + \frac{i_e^\mu}{A_\mu} + g_{\mu,\mu+1}(V_{\mu+1} - V_\mu) + g_{\mu,\mu-1}(V_{\mu-1} - V_\mu), \quad (3)$$

where  $\mu$  labels the compartments,  $i_e^\mu$  is the total electrode current flowing into the compartment  $\mu$ , and  $A_\mu$  is its surface area. The constant  $g_{\mu,\mu-1}$  determines the resistive coupling from neighboring compartment  $\mu - 1$  to compartment  $\mu$  and, for nonbranching cables, can be shown to be equal to  $g_{\mu,\mu-1} = a/(2r_L L^2)$  for a length  $L$  of each compartment. This defines a system of ordinary differential equations, which can be solved by generalized Euler methods.

## Problems

1. Numerically solve the cable equation (equation 1) when injecting a steady depolarizing current halfway along the cable. Consider that the membrane is passive, i.e.  $i_m = (V - E_L)/r_m$ , where  $r_m$  is the specific membrane resistance. Take  $r_m = 20 \text{ k}\Omega \text{ cm}^2$ ,  $r_L = 200 \Omega \text{ cm}$ ,  $c_m = 1 \mu\text{F}/\text{cm}^2$ , and cable radius  $a = 2 \mu\text{m}$ . Compare the solution to the analytical solution of the infinite cable.
2. Implement the Hodgkin-Huxley model of action potential propagation in the squid giant axon. Solve the partial differential equation using the Crank-Nicholson method (see e.g. Chapters 5.5, 5.6 and 6.6B from Dayan and Abbott, 2001). Take  $a = 238 \mu\text{m}$  and  $r_L = 35.4 \Omega \text{ cm}$ . Note that you will need to include your code with the project report!
3. Initiate an action potential on one end of the axon by injecting a current in the terminal compartment.
4. Determine the action potential propagation velocity as a function of the axon radius.
5. Initiate action potentials at both ends of the axon. Show that they annihilate when they collide.
6. Simulate action potential propagation in a myelinated axon (see e.g. Chapter 6.4 from Dayan and Abbott, 2001). Consider that the nodes of Ranvier (the unmyelinated spaces) are  $2 \mu\text{m}$  long and are located at  $1 \text{ mm}$  intervals along the axon. Also consider that myelin increases the resistance across the cell membrane by a factor of 5,000 and decreases the capacitance by a factor of 50.

## References

P Dayan, LF Abbott. Theoretical neuroscience, MIT Press 2001. Note the following if you have the first edition of the book. On page 226, equation 6.45: the symbol  $A_\mu$  in the equation for  $D_\mu$  denotes the surface area of compartment  $\mu$  of the cable, it is not the coefficient denoted by  $A_\mu$  that appears in the upper left equation and in equation 6.44.

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