

# REAL-TIME NEURAL SIGNAL FILTERING VIA HODGKIN-HUXLEY SIMULATION MODELS

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## ABSTRACT

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## 1 INTRODUCTION

## 2 MOTIVATION & RELEVANCE

In the field of biomedical engineering, extracting action potential timings from noisy extracellular recordings is essential in advancing brain-computer interfaces and neuroscience research. While spike detection itself is a signal processing task, the underlying signal can be described and approximated by the Hodgkin-Huxley (H-H) model. This set of non-linear ordinary differential equations models the ionic conductance changes that generate the action potential waveform. Our challenge is to adapt the H-H waveforms to mimic the noise present in real-life data.

Beyond common EEG readings or microelectrode arrays, neural signal filtering and spike detection is relevant to other closed-loop systems such as epileptic seizure prediction (?) or adaptive deep brain stimulation for Parkinson's disease (?). Advancements in filtering are further motivated by the advent of high-density neural probes which generate large data streams requiring efficient, accurate processing solutions (?). This project aims to develop a spike detection algorithm based on the H-H model to improve accuracy in low signal-to-noise ratio (SNR) environments.

## 3 SCOPE & FEASIBILITY

The scope of this project builds upon concepts from ESC103: Engineering Mathematics & Computation and MAT292: Ordinary Differential Equations. The work is divided into three primary phases: (1) generating synthetic neural data by solving the Hodgkin-Huxley equations, (2) processing this data with a digital filter, and (3) developing a spike detection algorithm.

### 3.1 PROJECT OBJECTIVES

The primary objectives of this project are:

1. **Data Generation:** To implement numerical solvers for the Hodgkin-Huxley (H-H) model to generate realistic synthetic action potential data.
2. **Signal Processing:** To design and apply a digital band-pass filter to isolate the spike waveform from the generated signal and added synthetic noise.
3. **Spike Detection:** To develop an algorithm that detects action potentials using an adaptive threshold, calculated from the estimated noise floor of the processed signal.
4. **Validation:** To qualitatively and quantitatively assess the performance of the detection algorithm on the noisy synthetic data.

### 3.2 PROJECT MILESTONES & TIMELINE

Although a more detailed outline of our project milestones and timeline can be found in A, several high-level milestones are listed below.

**Week 4:** Finish implementation of Euler's and Improved Euler's method solvers. Also complete first iteration of noise generation algorithm.

**Week 6:** Finish implementation of Runge-Kutta method solver(s), and complete first iteration of band-pass filter.

**Week 8:** Refine band-pass filter, noise generation algorithm, and numerical methods.

**Week 10:** Discuss and evaluate results in the final report.

**Week 11/12:** Buffer time in case aforementioned tasks take longer than initially anticipated.

## 4 TECHNICAL BACKGROUND

Understanding the biological model of neuron signals is imperative to acknowledging the abstractions and simplifications made to obtain the H-H equations. In this section, we overview the basics behind action potential in the brain, and biochemical processes that occur to produce these phenomena. Additionally, we outline the handful of key equations we intend to use from the H-H model, and discuss the theory behind the numerical methods we have chosen.

### 4.1 THE BIOLOGICAL BASIS: ION CHANNELS AND CURRENTS

Electrophysiology—the study of electrical properties of cells—provides the basis for cell communication via current signals. This supports not only the neural signalling process, but also fundamental processes like muscle contraction, nutrient transport, and cellular homeostasis. These current signals work tangentially with ion channels—pores discovered by Hodgkin and Huxley that use selective permeability to regulate intramolecular transport across lipid bilayers. Ion channels occur when current flows across a cell membrane. This transport occurs bi-directionally, its flux reversing as the Nernst potential is reached (zero-current state). Ion channel movement can be measured via the ionic currents that pass through, this measurement is based on Ohm's Law (CITE HERE, RUBAIY).

#### 4.1.1 THE CELL MEMBRANE AND RESTING POTENTIALS

Action potential is the change in membrane electric potential that creates ion channels (CITE, RUBAIY). The cell electrical state is measured as  $V_m$ , the potential difference between the cytoplasm and external environment (CITE, CERVERA). In their resting state, cell membranes possess a net negative charge (resting potential), preventing ion channel creation. Depolarization occurs when a membrane changes from a net negative charge to a net positive charge, opening an ion channel. Repolarization occurs when the membrane regains its negative charge, closing the ion channel (CITE, RUBAIY). This electrical bi-stability of depolarization to repolarization is an important characteristic of neural cells (CITE, RUBAIY).

#### 4.1.2 VOLTAGE-GATED ION CHANNELS

#### 4.1.3 SODIUM-POTASSIUM PUMP

### 4.2 MODELING NEURONS AS ELECTRICAL CIRCUITS

#### 4.2.1 THE PARALLEL CONDUCTANCE MODEL

#### 4.2.2 CIRCUIT ANALOGS: CAPACITORS, RESISTORS, AND BATTERIES

### 4.3 HODGKIN-HUXLEY EQUATIONS

The Hodgkin-Huxley model is a system of nonlinear differential equations that represent action potentials movement through nerve cells. Hodgkin and Huxley's experiments revealed three types of ion current: sodium, potassium, and a leak current. To discern equations that model the electrical properties of a cell during the action potential, Hodgkin and Huxley represented the transfer of ions across the cell membrane as a circuit. These equations laid the basis for understanding ion channel kinetics and nerve cell excitability.

#### 4.3.1 MODEL COMPONENTS AND EQUATIONS

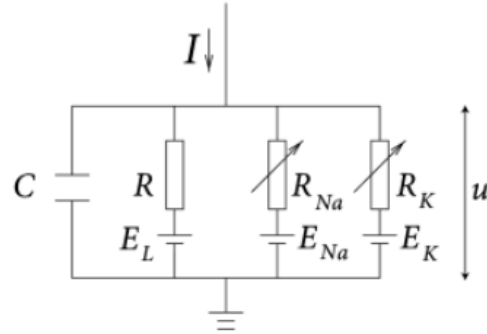


Figure 1: Circuit representation of ion transfer across the cell membrane

The circuit model represents each component of ion transfer within a nerve cell as an electrical component: the ion pumps and the leak channel are represented as resistors; the difference in ion concentration for each ion is represented by a battery; the lipid bilayer is represented by a capacitor. These components are combined according to the circuit in Figure 1, to create the equation for current across the cell membrane:

$$I = C_M \frac{dV}{dt} + \bar{g}_K n^4 (V - V_K) + \bar{g}_{Na} m^3 h (V - V_{Na}) + \bar{g}_l (V - V_l) \quad (1)$$

Relating Equation 1 to Figure 1,  $V$  is analogous to  $u$ , and  $V_x$  is analogous to  $E_x$  (where  $x$  represents Na, K, or l). Discerned from the resistance values,  $g_x$  (where

$x$  is Na, K, or I) represents the maximum conductance of the channel.

#### 4.3.2 GATING VARIABLES AND ACTIVATION/INACTIVATION

Within the equations there are three gating variables ( $m$ ,  $n$ , and  $h$ ) that represent the probability that a channel will be open at a given moment in time. The  $Na^+$  channels are controlled by  $m$  and  $h$ , where  $m$  describes the activation of the channel and  $h$  is its inactivation. Similarly,  $n$  describes the activation of the  $K^+$  channels.

After observing delayed-onset kinetics in their experiments, Hodgkins and Huxley theorized that a specific amount of charged particles would need to move under influence of the membrane potential to allow  $Na^+$  or  $K^+$  particles to move. They found that  $n^4$ ,  $m^3$ , and  $h$  provided a good fit. From this data, differential equations were developed to model the gating variables for a range of potentials. These equations take the form:

$$\dot{x} = -\frac{1}{\tau_x(u)}[x - x_0(u)] \quad (2)$$

To interpret Equation 2, let  $\dot{x} = dx/dt$  where  $x$  stands for  $m$ ,  $n$ , or  $h$ .  $\tau_0(u)$  and  $x_0(u)$  are conditional variables dependent on the initial membrane potential.

#### 4.4 NUMERICAL METHODS

Oftentimes in real-world modelling, differential equations do not have an algebraic solution. This does not impede our ability to visualize or obtain results as we can instead turn to numerical methods to approximate the solution. Within the field of numerical methods and computation, there are numerous methods to approximate a solution, however we will focus on three main methods: Euler's Method, Improved Euler's Method, and Runge-Kutta Methods.

##### 4.4.1 EULER'S METHOD

Intuitively, Euler's Method solves the initial value problem by linking tangent lines for a finite number of time steps of size  $\Delta t$ . Given a differential equation and initial condition:

$$\frac{dy}{dt} = f(t, y) \quad \text{where} \quad y(t_0) = y_0$$

we can approximate the tangent line of the solution at the initial point  $(t_0, y_0)$  by:

$$y = y_0 + f(t_0, y_0)(t - t_0)$$

We approximate the solution at some  $y_i$  by using the tangent line from the previous point (starting at  $t_0$ ) to

obtain more points. As we increase the number of time steps and consequently decrease the size of  $\Delta t$ , we can improve our approximation of the solution curve. Thus, we can generalize the approximation of the solution  $\phi(t)$  with the following equation:

$$y_{n+1} = y_n + f(t_n, y_n) \cdot (t_{n+1} - t_n)$$

##### 4.4.2 IMPROVED EULER'S METHOD

Though Euler's Method can give us a relatively concrete idea of the behaviour of the solution, it requires a large number of time steps to achieve accurate results. Thus, many such methods aim to improve upon this base including Improved Euler's Method. Intuitively, this method simply builds upon Euler's Method by considering the derivative at the subsequent point and computing the average. We can define the general formula for the Improved Euler's Method with the following:

$$y_{n+1} = y_n + \frac{f_n + f(t_n + h, y_n + hf_n)}{2}h$$

such that

$$h = t_{n+1} - t_n \text{ and } f_n = f(t_n, y_n)$$

##### 4.4.3 RUNGE-KUTTA METHODS

Euler's Method and Improved Euler's Method are technically considered Runge-Kutta class of numerical methods; however, in this section we will discuss the original methods developed by Runge and Kutta. This method is classified as a fourth-order, four-stage Runge Kutta Method, but is commonly referred to as "the" Runge-Kutta Method. Essentially, this method builds upon the two aforementioned methods by computing a weighted average of values at different points along the time step  $\Delta t$ . The general form for the approximation of the solution is given by:

$$y_{n+1} = y_n + h \cdot \frac{k_{n1} + 2k_{n2} + 2k_{n3} + k_{n4}}{6}$$

where we define each  $k_{ni}$  as the following:

$$\begin{aligned} k_{n1} &= f(t_n, y_n) \\ k_{n2} &= f\left(t_n + \frac{1}{2}h, y_n + \frac{1}{2}hk_{n1}\right) \\ k_{n3} &= f\left(t_n + \frac{1}{2}h, y_n + \frac{1}{2}hk_{n2}\right) \\ k_{n4} &= f(t_n + h, y_n + hk_{n3}) \end{aligned}$$

## 5 CONCLUSION

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