Module1 Ex2 09 off

March 14, 2024

1 Lab Session #2

1.1 Computational Neurophysiology [E010620A]

1.1.1 Dept of Electronics and Informatics (VUB) and Dept of Information Technology (UGent)

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1.1.2 General Introduction

In all the practical sessions of this course we will use python 3 and jupyter notebooks. Please install anaconda on your computer and after installation you can open jupyter notebook by typing "jupyter notebook" in the command line. Your browser will open a search directory, which you can use to browse to and open the exercise. Alternatively, you can use jupyter-lab.

Deadline: 2 weeks after lecture

The lab sessions consist of a jupyter notebook in which the different steps are described and explained, together with the tasks that students are asked to complete.

This practical is based upon the freely available python exercise: https://neuronaldynamics-exercises.readthedocs.io/en/latest/exercises/adex-model.html

1.1.3 Context and Goals

This second lab session is focused on the Adaptive Exponential Integrate-and-Fire model. The students are asked to implement the equations as seen in the lecture (and repeated here) and describe what they see in different simulations.

Whereas most of coding can be done without the BRIAN package, it can be a useful tool to check your own results.

2 Questions

2.1 1 AdEx Integrate-and-Fire model

In this first part, we will code and develop the Adaptive exponential integrate-and-fire model, without the use of the BRIAN library. To complete this task, start from the theoretical chapter https://neuronaldynamics.epfl.ch/online/Ch6.S1.html and the following equations:

$$\begin{split} \tau_m \frac{\mathrm{d}u}{\mathrm{d}t} &= -(u - u_{\mathrm{rest}}) + \Delta_T \exp\left(\frac{u - \theta_{\mathrm{rh}}}{\Delta_T}\right) - Rw + RI(t) \\ \tau_w \frac{\mathrm{d}w}{\mathrm{d}t} &= a(u - u_{\mathrm{rest}}) - w + b\tau_w \sum_{tf} \delta(t - t^f) \end{split}$$

The following constants can be used for the model parameters. Note that the BRIAN package uses units. Whereas this is not required for your own coding, make sure that the units match!

• Import these modules

2.1.1 Q1 Generate input current

Q1a The first step is to generate the input current I(t). For this we create a step function of length 370 ms. The input current is 0 μ A at t=0 and steps to 1 μ A at t=20ms. The input current is reset to 0 μ A at t=200ms. Create and plot I_input in function of t and make sure that the time step is 0.01 ms. This timestep corresponds to the integration step when we will solve the differential equations and can remain constant for the purpose of this practical.

Q1b Create a function that outputs u(t), w(t), DeltaU(t) and DeltaW(t) in function of the initial values of u and w (u_0,w_0) and the input current I_input(t). Please also print the time point whenever an action potential is being fired.

Q1c Test this function with the input current that you have defined previously but with an amplitude of $68 \mu A$ and create five plots below each other: - I(t) - u(t) - w(t) - DeltaU(t) - DeltaU(t)

The initial value of u is u_rest (-70 mV), the initial value of w can be set to zero.

Q1d Describe the evolution between subsequent action potentials. Plot the evolution of these intervals. What do you notice?

• Fill in answer here

2.2 2 BRIAN Library - I&F models

Here we will implement the non-adaptive and adaptive exponential integrate-and-fire model through the BRIAN package. First things first, the non-adaptive I&F model: - Again we need to create an input current. Within the BRIAN package the same input profile as before can be easily calculated with the input_factory.get_step_current() function - Next, we need to simulate the model. This can be done through the exp_IF() function. Which are the default values of this model? - Finally, we plot our output with the plot_tools.plot_voltage_and_current_traces() tool.

2.2.1 Q2.1 Exponential Integrate and Fire

Apply the suggested functions to simulate the behaviour of a firing neuron when the exponential integrate and fire model is used. 1. Apply a step input current of amplitude 0.9 nA that starts at t = 20 ms and ends at t = 150 ms 2. Simulate what happens for 200 ms

How many spikes do you get?

• Fill in answer here

2.2.2 Q2.2 Adaptive Exponential I&F - BRIAN

What happens when you substitute the non-adaptive by the adaptive exponential model? You can use the simulate_AdEx_neuron function.

- 1. Apply an input current of amplitude 90 pA that starts at t = 50 ms and ends at t = 150 ms.
- 2. Simulate what happens for 350 ms using simulate_AdEx_neuron

How many spikes are you getting now?

• Fill in answer here

2.2.3 Q2.3 Characteristics

Which are the characteristics of the AdEx model? How many spikes do you observe? Describe the firing pattern.

• Fill in answer here

2.3 3 Firing Pattern

2.3.1 Q3 Simulate all patterns

By changing the parameters in the function AdEx.simulate_AdEx_neuron(), you can simulate different firing patterns. Create tonic, adapting, initial burst, bursting, irregular, transient and delayed firing patterns. Table 6.1 provides a starting point.

Simulate your model for 350 ms and use a step current of 67 pA starting at t = 50 to t = 250.

• Fill in answer here

2.4 4 Phase plane and Nullclines

In this section, you will acquire some intuition on shape of nullclines by plotting and answering the following questions.

• Import these modules

2.4.1 Q4.1 Run AdEx

Plot the u and w nullclines of the AdEx model 1. How do the nullclines change with respect to a? 2. How do the nullclines change if a constant current I(t) = c is applied? 3. What is the interpretation of parameter b? 4. How do flow arrows change as tau_w gets bigger?

For this plot, you won't need the BRIAN library, but you can use functions that are available through numpy. You will need to create a grid of u, w values through np.meshgrid. Next, for each point of this grid, you will have to evaluate the time-derivative (Formulas 6.3 and 6.4). Finally, you will have to calculate the null-clines and plot everything together on a single plot. For the plotting of the arrows, you can have a look at the np.quiver function.

• Fill in answer here

2.4.2 Q4.2 Predict firing pattern

Can you predict what would be the firing pattern if the value 'a' is small (in the order of 0.01 nS)? To do so, consider the following 2 conditions:

A large jump b and a large time scale tau_w. A small jump b and a small time scale tau_w. Try to simulate the above conditions, to see if your predictions were correct.

• Fill in answer here

3 Answers

3.1 1 AdEx Integrate-and-Fire model

3.1.1 Import

```
[2]: # Here add all the libraries and modules that are needed throughout the notebook
import math
import numpy as np
import matplotlib.pyplot as plt
import brian2 as b2

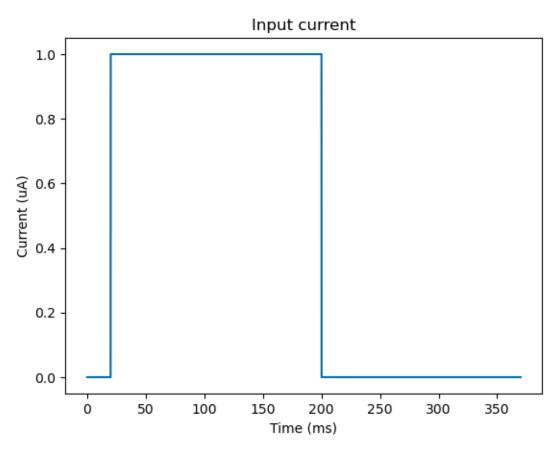
# Make your graphs color blind friendly
plt.style.use('tableau-colorblind10')
```

3.1.2 A1 Generate input current

• Go back to Q1

```
# create I_input
def I_input(t):
    return np.where(t < t1, 0, np.where(t < t2, I0, 0))

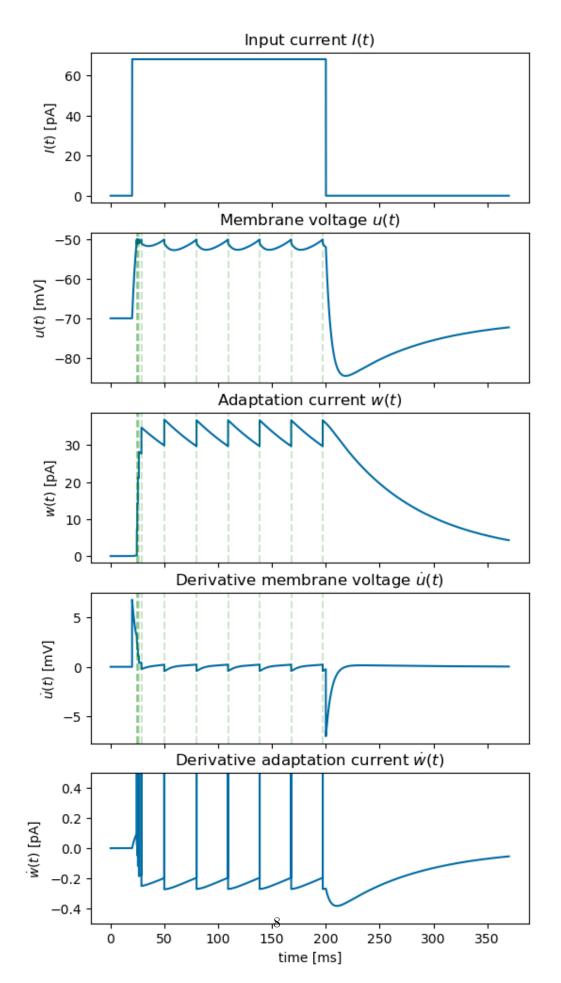
# plot I_input
t = np.arange(0, T, dt)
plt.plot(t, I_input(t))
plt.xlabel('Time (ms)')
plt.ylabel('Current (uA)')
plt.title('Input current')
plt.show()</pre>
```

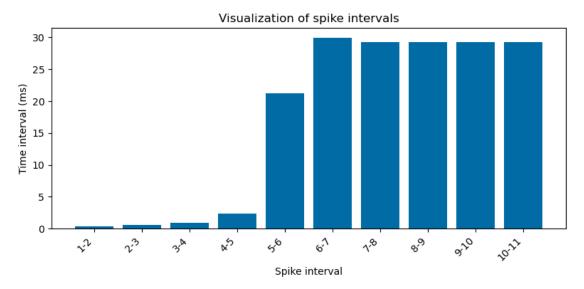


```
# parameters
  N = int(T_sim / dt)
                                      # number of time steps
  R_m = params['R_m']
                                       # GOhm
  u_rest = params['u_rest']
                                       # mV
  u_reset = params['u_reset']
                                       # mV
  v_rheobase = params['v_rheobase'] # mV
  delta_T = params['delta_T']
                                      # ms
  a = params['a']
                                       \# nS
  tau_w = params['tau_w']
                                       # ms
  tau_m = params['tau_m']
                                       # ms
  b = params['b']
                                       # pA
  spike_index = []
  # vector objects
  u, w = np.zeros(N), np.zeros(N)
  t = np.arange(0, T_sim, dt)
  I = I_input(t)
  # set the initial coonditions
  \mathbf{u}[0] = \mathbf{u}_0
  w[0] = w_0
  # update the model
  for k in range(1, N):
       # update u, w
      u[k] = u[k-1] + dt/tau_m * (-(u[k-1] - u_rest) + delta_T*np.exp((u[k-1]_u))
\rightarrow v_rheobase) / delta_T) - R_m*w[k-1] + R_m*I[k-1])
       w[k] = w[k-1] + dt/tau_w * (a*(u[k-1] - u_rest) - w[k-1])
       # check if firing occurs
      if u[k-1] > v_rheobase:
           u[k] = u_reset
           w[k] += b
           spike_index.append(k)
  # calculate the derivatives
  delta_us = (u_rest - u + delta_T * np.exp((u-v_rheobase)/delta_T) + R_m*(I_U
⊶ w)) / tau_m
  delta_ws = (a*(u - u_rest) - w) / tau_w
  delta_ws[spike_index] += b
  # construct spike_times
  spike_times = np.array(spike_index) * dt
  return u, w, delta_us, delta_ws, spike_times
```

```
params_tonic = {'tau_m':20, 'a':0, 'tau_w':30, 'b':60, 'u_reset':-55, 'u_rest':__
     ↔-70, 'delta_T':2, 'R_m':0.500, 'v_rheobase':-50}
    params_transient = {'tau_m':10, 'a':1.0, 'tau_w':100, 'b':10, 'u_reset':-60,__
     params bursting = {'tau m':5.0, 'a':-0.5, 'tau w':100, 'b':7.0, 'u reset':-46,,,
     params irragular = {'tau m':9.9, 'a':-0.5, 'tau w':100, 'b':7.0, 'u reset':-46,
     u, w, delta_us, delta_ws, spike_times = adex(-70, 0, lambda t: 68*I_input(t)) #__
     \hookrightarrow I is in pA
    print('the timepoints when a spike has occured are: \n {} ms'.
     →format(spike_times))
   the timepoints when a spike has occured are:
                 25.37 26.3
    [ 24.38 24.8
                             28.67 49.9
                                        79.87 109.15 138.47 167.78
    197.09] ms
fig, axs = plt.subplots(5, 1, figsize = (6,12), sharex = True)
    axs[0].plot(t, 68*I input(t))
    axs[1].plot(t, u)
    axs[2].plot(t, w)
    axs[3].plot(t, delta_us)
    axs[4].plot(t, delta_ws)
    # plot the spike times on each polot
    for ax in axs[1:-1]:
       for spike_time in spike_times:
          ax.axvline(spike_time, color = 'g', linestyle = '--', alpha = 0.2)
    axs[0].set title('Input current $I(t)$')
    axs[1].set_title('Membrane voltage $u(t)$')
    axs[2].set title('Adaptation current $w(t)$')
    axs[3].set title('Derivative membrane voltage $\dot{u}(t)$')
    axs[4].set title('Derivative adaptation current $\dot{w}(t)$')
    axs[0].set_ylabel('$I(t)$ [pA]')
    axs[1].set_ylabel('$u(t)$ [mV]')
    axs[2].set_ylabel('$w(t)$ [pA]')
    axs[3].set_ylabel(r'$\dot{u}(t)$ [mV]')
    axs[4].set_ylabel(r'$\dot{w}(t)$ [pA]')
    axs[4].set_xlabel('time [ms]')
    axs[4].set_ylim((-0.5, 0.5))
```

plt.show()





A1 conclusion:

For the simulation above, parameters for *Initial Bursting* (Table 6.1) were used.

0. Mathematics of the AdEx IF model

The memebrane potential u in the Adaptive Exponential Integrate-and-Fire (AdEx IF) model, is

described by the differential equations:

$$\begin{split} \tau_m \frac{\mathrm{d}u}{\mathrm{d}t} &= -(u - u_{\mathrm{rest}}) + \Delta_T \exp\left(\frac{u - \theta_{\mathrm{rh}}}{\Delta_T}\right) - Rw + RI(t) \\ \tau_w \frac{\mathrm{d}w}{\mathrm{d}t} &= a(u - u_{\mathrm{rest}}) - w + b\tau_w \sum_{tf} \delta(t - t^f). \end{split}$$

Where w(t) is the adaptation current, that is coupled to the membrane potential by a. When the membrane potential reaches the threshold $\theta_{\rm rh}$ at $t=t^f$, a spike is generated and the membrane potential is reset to $u_{\rm reset}$. At $t=t^f$, the adaptation current is increased with b. After firing, the integration of the membrane potential is again described by the AdEx differential equations. As will be discussed in other conclusions the parameters a and b are the source of the subtreshold adaptation.

1. Describe the evolution between subsequent action potentials.

Just after the generation of a spike 2 , the membrane potential is reset to $u_{\rm reset}$ and the adaptation current is increased with b^2 . While the constant input current still persists, the membrane potential starts integrating again $\dot{u}>0$ and the adaptation current decreases $\dot{w}<0^3$. When the membrane potential reaches the threshold again, this procedure starts again.

2. Plot the evultion of these intervals.

The evolution of the interspiking intervals is plotted in a bar chart (see above).

3. What do you notice?

One can notice that the spiking pattern consists of a *transient* phase just after the onset of the input current and a *steady-state* phase: the neuron starts adapting and (in this case) starts firing in a periodic manner. In the simultaned model above, the transient phase is characterized by an initial burst of firing (short interspike intervals in the bar chart) and the steady state phase exists after 6 spikes (equal interspike intervals in the bar chart).

- ¹ Note that in the plot of u(t) above the 'spikes' are not plotted. The AdEx IF model assumes there is no information in the shape of the spike, and thus plotting the spike itself is thus irrelevant.
- ² You can see the jumps in the plots of u(t) and w(t).
- ³ Note that in the plots above there are discontinuities in $\dot{u}(t)$ and $\dot{w}(t)$ due to the incontinuities in u(t) and w(t) at $t=t^f$. This is the result of a computation error because at the $t=t^f$, $\dot{u}(t)$ and $\dot{w}(t)$ are not defined! So, if I discuss $\dot{u}(t)$ or $\dot{w}(t)$, I only consider the values where they exist: $t \in \text{dom}(\dot{u})$ or $t \in \text{dom}(\dot{w})$ respectively.

3.2 2 BRIAN Library - I&F models

3.2.1 Import

```
[8]: %matplotlib inline
import brian2 as b2
import neurodynex3.exponential_integrate_fire.exp_IF as exp_IF
from neurodynex3.tools import plot_tools, input_factory
from neurodynex3.adex_model import AdEx
```

3.2.2 A2.1 Exponential Integrate and Fire

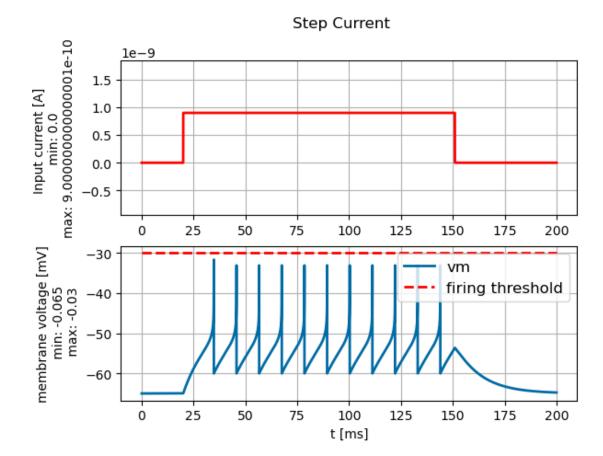
• Go back to Q2.1

```
# set_current
    t_start, t_end = 20, 150
    A = 0.9
    I_input = input_factory.get_step_current(t_start, t_end, b2.ms, A*b2.nA)
    # simulation of the model (What are the default values of this model?)
    T \sin = 200
    state monitor, spike monitor = exp IF.
     simulate_exponential_IF_neuron(I_stim=I_input, simulation_time = T_sim * b2.
     oms)
    n_spikes = spike_monitor.count[0]
    print("n_spikes: {}".format(n_spikes))
    # plot the output of the model
    plot_tools.plot_voltage_and_current_traces(state_monitor, I_input, title = u
      → 'Step Current', firing threshold=exp IF.FIRING THRESHOLD v spike)
    WARNING
              Cannot use Cython, a test compilation failed: Cython is not available
    (ImportError) [brian2.codegen.runtime.cython rt.cython rt.failed compile test]
              Cannot use compiled code, falling back to the numpy code generation
    target. Note that this will likely be slower than using compiled code. Set the
    code generation to numpy manually to avoid this message:
    prefs.codegen.target = "numpy" [brian2.devices.device.codegen_fallback]
    n spikes: 11
[9]: (<AxesSubplot:ylabel='Input current [A] \n min: 0.0 \nmax:
```

<AxesSubplot:xlabel='t [ms]', ylabel='membrane voltage [mV]\n min: -0.065\n</pre>

9.00000000000001e-10'>,

max: -0.03'>)



A2.1 conclusion:

In the simulation above, one can observe a that a periodic firing pattern occurs. I.e. there is not transient phase. In total 11 spikes were generated.

3.2.3 A2.2 Adaptive Exponential I&F - BRIAN

• Go back to Q2.2

[10]: # getting the default values via the help command and looking in the code for static variables (e.g. V_RESET) help(exp_IF)

Help on module neurodynex3.exponential_integrate_fire.exp_IF in neurodynex3.exponential_integrate_fire:

NAME

neurodynex3.exponential_integrate_fire.exp_IF

DESCRIPTION

Exponential Integrate-and-Fire model.

See Neuronal Dynamics, `Chapter 5 Section 2

```
<http://neuronaldynamics.epfl.ch/online/Ch5.S2.html>`_
FUNCTIONS
   getting_started()
       A simple example
    simulate exponential IF neuron(tau=12. * msecond, R=20. * Mohm, v rest=-65.
* mvolt, v_reset=-60. * mvolt, v_rheobase=-55. * mvolt, v_spike=-30. * mvolt,
delta_T=2. * mvolt, I_stim=<br/>brian2.input.timedarray.TimedArray object at
0x7f872d876730>, simulation_time=200. * msecond)
        Implements the dynamics of the exponential Integrate-and-fire model
        Args:
            tau (Quantity): Membrane time constant
            R (Quantity): Membrane resistance
            v_rest (Quantity): Resting potential
            v_reset (Quantity): Reset value (vm after spike)
            v_rheobase (Quantity): Rheobase threshold
            v_spike (Quantity) : voltage threshold for the spike condition
            delta_T (Quantity): Sharpness of the exponential term
            I stim (TimedArray): Input current
            simulation time (Quantity): Duration for which the model is
simulated
        Returns:
            (voltage_monitor, spike_monitor):
            A b2.StateMonitor for the variable "v" and a b2.SpikeMonitor
DATA
   FIRING_THRESHOLD_v_spike = -30. * mvolt
   MEMBRANE_RESISTANCE_R = 20. * Mohm
   MEMBRANE_TIME_SCALE_tau = 12. * msecond
   RHEOBASE_THRESHOLD_v_rh = -55. * mvolt
   SHARPNESS_delta_T = 2. * mvolt
    V RESET = -60. * mvolt
    V_REST = -65. * mvolt
FILE.
    /Users/constantijncoppers/anaconda3/envs/cn/lib/python3.8/site-
packages/neurodynex3/exponential_integrate_fire/exp_IF.py
```

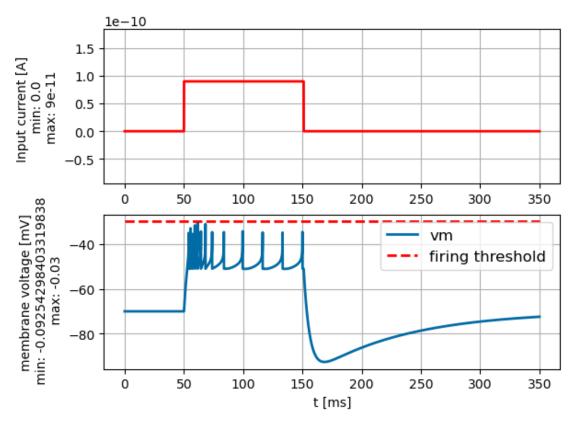
from neurodynex3.tools import plot_tools, input_factory

from neurodynex3.adex_model import AdEx

```
# set_current
t_start, t_end = 50, 150
A = 90
I_input = input_factory.get_step_current(t_start, t_end, b2.ms, A*b2.pA)

# simulation of the model (What are the default values of this model?)
T_sim = 350
state_monitor, spike_monitor = AdEx.simulate_AdEx_neuron(I_stim = I_input,_u_simulation_time = T_sim * b2.ms)
n_spikes = spike_monitor.count[0]
print("n_spikes: {}".format(n_spikes))

# plot the output of the model
plot_tools.plot_voltage_and_current_traces(state_monitor, I_input, title =_u_s'Step Current', firing_threshold=exp_IF.FIRING_THRESHOLD_v_spike)
```



3.2.4 A2.3 Characteristics

• Go back to Q2.3

A2.2 and A2.3 answer:

1. Characteristics of the AdEx model

Lets break down the characteristics of AdEx by explaining its name:

1. **Exponential:** The model incorporates a nonlinear function f(u) inspired by empirical data. This function comprises a linear leak term and an exponential activation term:

$$f(u) = \underbrace{-\left(u - u_{\text{rest}}\right)}_{\text{linear leak}} + \underbrace{\Delta_T \exp\left(\frac{u - \theta_{\text{rh}}}{\Delta_T}\right)}_{\text{exponential activation term}}.$$

 Δ_T is the sharpness of action potential initiation and $\theta_{\rm rh}$ the rheobase threshold (see further on for interpretation).

- 2. Adaptation: AdEx features a single adaptation current w, which evolves with the time constant τ_w and according to a differential equation (6.4) involving parameters a, and b. The parameters a and b relate to the dynamics of ion channels. The parameter a governs the coupling between the adaptation current w and the membrane potential u, while b influences the increase in the current at spike times. These parameters shape the spiking pattern of the neuron, hence characterizing the 'Adaptive' aspect of the model.
- 3. Integrate-and-fire dynamics: The model assumes that the information encoded in neuronal activity primarily resides in the spike pattern rather than the precise shape action potentials. Consequently, when the membrane potential reaches a certain threshold $\theta_{\rm rh}$, it triggers a spike, after which the membrane potential is reset and the adaptation current increased by b. This feature underscores the 'Integrate-and-Fire' nature of the AdEx model. Altough f is nonlinear it still remains simple enough to predict the firing pattern in response to a constant input current.

2. Observed spikes

In total 13, spikes were observed. However, the duration of the current in Q2.1 is longer so you can not absolutely compare them. Though you can state that there will be more spikes evoked in Q2.2 model since the current duration there was shorter and lead to a higher number of evoked spikes.

3. Firing pattern

In contrast to the firing pattern observed in the simulation of Q2.1, where the model exhibits a regular spiking pattern from the beginning, the simulation in Q2.2 initially displays a burst of spikes followed by a transition to a regular spiking pattern. This initial burst of spikes characterizes the transient phase of the firing pattern, which eventually evolves into a steady-state phase of regular spiking.

The difference in firing patterns between Q2.1 and Q2.2 arises from the inclusion of adaptation mechanisms in Q2.2. Specifically, the adaptation mechanism modeled in Q2.2 contributes to the transient phase, leading to the observed burst of spikes. In contrast, Q2.1 does not incorporate any adaptation mechanism, resulting in a periodic spiking pattern without a transient phase.

Therefore, the presence of the transient phase in Q2.2, driven by adaptation dynamics, leads to a higher number of spikes compared to Q2.1, where no such transient phase is modeled.

3.3 3 Firing Pattern

3.3.1 A3 Simulate all patterns

• Go back to Q3

```
[12]: # fixed parameters #
# step current
t_start, t_end = 50, 250
A = 67
I_input = input_factory.get_step_current(t_start, t_end, b2.ms, A*b2.pA)

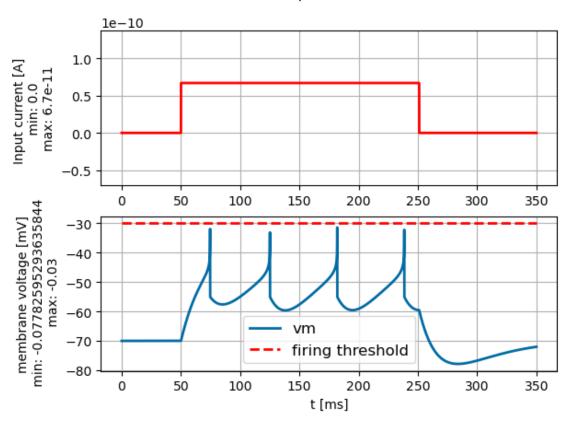
# simulation parameters
u_rest = - 70 * b2.mV
delta_T = 2 * b2.mV
v_rheobase = - 50 * b2.mV
R_m = 500 * b2.Mohm

# check how to define the parameters
# help(AdEx)
```

```
[13]: #Tonic
      # parameters
      u_reset = -55.0 * b2.mV
      tau m = 20.0 * b2.ms
      tau_w = 30.0 * b2.ms
      a = 0.0 * b2.nS
      b = 60 * b2.pA
      # simulate the model
      state_monitor, spike_monitor = AdEx.simulate_AdEx_neuron(I_stim = I_input,_
       ⇒simulation_time = T_sim * b2.ms,
                                                                v_reset = u_reset,_

¬v_rest = u_rest, v_rheobase = v_rheobase, delta_T = delta_T,
                                                                tau_m = tau_m, tau_w =
       ⇔tau_w,
                                                                a = a, b = b,
                                                                R = R_m)
      n_spikes = spike_monitor.count[0]
      print("n_spikes: {}".format(n_spikes))
      # plot the output of the model
```

```
plot_tools.plot_voltage_and_current_traces(state_monitor, I_input, title = \_ \times \'Step Current', firing_threshold=exp_IF.FIRING_THRESHOLD_v_spike)
```

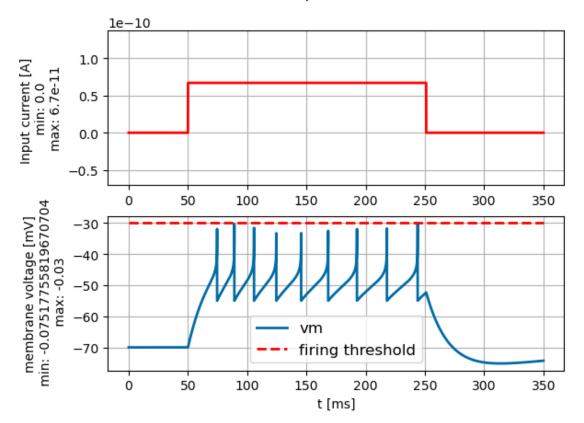


```
[14]: #Adapting
    # parameters
    u_reset = -55.0 * b2.mV

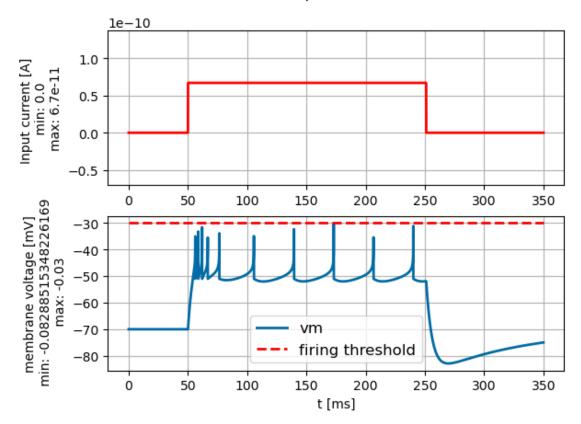
tau_m = 20.0 * b2.ms
tau_w = 100.0 * b2.ms

a = 0.0 * b2.nS
b = 5.0 * b2.pA

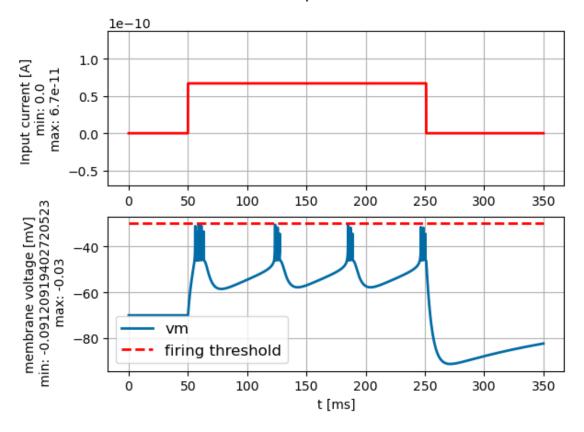
# simulate the model
```



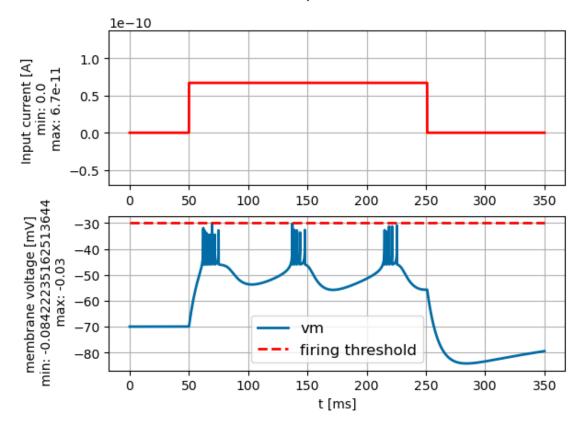
```
[15]: #Initial burst
     # parameters
     u_reset = -51.0 * b2.mV
     tau_m = 5.0 * b2.ms
     tau_w = 100.0 * b2.ms
     a = 0.5 * b2.nS
     b = 7.0 * b2.pA
     # simulate the model
     state_monitor, spike_monitor = AdEx.simulate_AdEx_neuron(I_stim = I_input,__
      ⇒simulation_time = T_sim * b2.ms,
                                                         v_reset = u_reset,__
      →v_rest = u_rest, v_rheobase = v_rheobase, delta_T = delta_T,
                                                         tau_m = tau_m, tau_w =
      ⇔tau_w,
                                                         a = a, b = b,
                                                         R = R_m)
     n_spikes = spike_monitor.count[0]
     print("n_spikes: {}".format(n_spikes))
     # plot the output of the model
     plot_tools.plot_voltage_and_current_traces(state_monitor, I_input, title =__
```



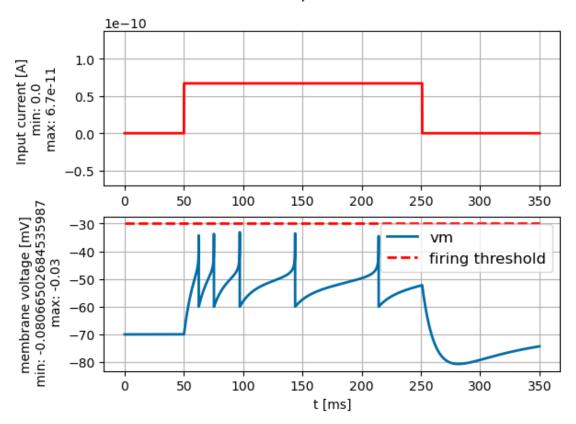
```
[16]: #Bursting
      # parameters
      u_reset = -46.0 * b2.mV
      tau_m = 5.0 * b2.ms
      tau_w = 100.0 * b2.ms
      a = -0.5 * b2.nS
      b = 7.0 * b2.pA
      # simulate the model
      state_monitor, spike_monitor = AdEx.simulate_AdEx_neuron(I_stim = I_input,__
       ⇔simulation_time = T_sim * b2.ms,
                                                               v_reset = u_reset,_
       ⇒v_rest = u_rest, v_rheobase = v_rheobase, delta_T = delta_T,
                                                               tau_m = tau_m, tau_w =
       ⇔tau_w,
                                                               a = a, b = b,
                                                               R = R m
```



```
a = -0.5 * b2.nS
b = 7.0 * b2.pA
# simulate the model
state_monitor, spike_monitor = AdEx.simulate_AdEx_neuron(I_stim = I_input,__
⇒simulation_time = T_sim * b2.ms,
                                                  v_reset = u_reset,_
→v_rest = u_rest, v_rheobase = v_rheobase, delta_T = delta_T,
                                                  tau_m = tau_m, tau_w =
→tau_w,
                                                  a = a, b = b,
                                                  R = R_m)
n_spikes = spike_monitor.count[0]
print("n_spikes: {}".format(n_spikes))
# plot the output of the model
plot_tools.plot_voltage_and_current_traces(state_monitor, I_input, title = ___
```



```
[18]: #Transient
      # parameters
      u_reset = -60.0 * b2.mV
      tau_m = 10.0 * b2.ms
      tau_w = 100.0 * b2.ms
      a = 1.0 * b2.nS
      b = 10.0 * b2.pA
      # simulate the model
      state_monitor, spike_monitor = AdEx.simulate_AdEx_neuron(I_stim = I_input,__
       →simulation_time = T_sim * b2.ms,
                                                               v_reset = u_reset,_
       ⇒v_rest = u_rest, v_rheobase = v_rheobase, delta_T = delta_T,
                                                               tau_m = tau_m, tau_w =
       ⇔tau_w,
                                                               a = a, b = b,
                                                               R = R m
```

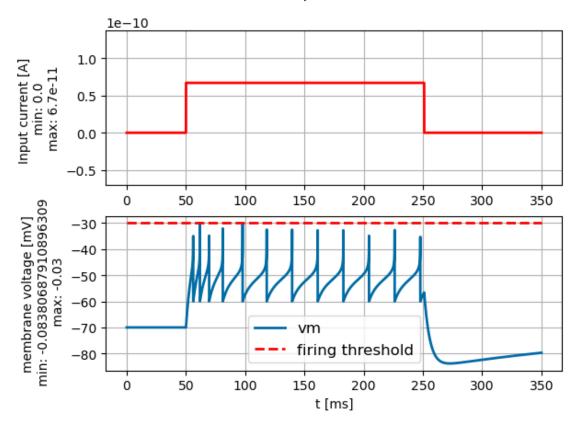


```
[19]: #Delayed
    # parameters
u_reset = -60.0 * b2.mV

tau_m = 5.0 * b2.ms
tau_w = 100.0 * b2.ms
```

```
a = -1.0 * b2.nS
b = 10.0 * b2.pA
# simulate the model
state_monitor, spike_monitor = AdEx.simulate_AdEx_neuron(I_stim = I_input,__
⇒simulation_time = T_sim * b2.ms,
                                                         v_reset = u_reset,_
 →v_rest = u_rest, v_rheobase = v_rheobase, delta_T = delta_T,
                                                         tau_m = tau_m, tau_w =
 →tau_w,
                                                         a = a, b = b,
                                                         R = R_m)
n_spikes = spike_monitor.count[0]
print("n_spikes: {}".format(n_spikes))
# plot the output of the model
plot_tools.plot_voltage_and_current_traces(state_monitor, I_input, title = ___ 
 Step Current', firing_threshold = exp_IF.FIRING_THRESHOLD_v_spike)
```

Step Current



3.4 4 Phase plane and Nullclines

3.4.1 Import

```
[20]: %matplotlib inline
import brian2 as b2
from neurodynex3.adex_model import AdEx
from neurodynex3.tools import plot_tools, input_factory
```

3.4.2 A4.1 Run AdEx

• Go back to Q4.1

```
[21]: # parameters

u_rest = -70  # mV

delta_T = 2  # mV

v_rheobase = -50  # mV

R_m = 0.500  #GOhm

u_reset = -51.0  # mV
```

```
# ms
tau m = 5.0
tau_w = 100.0
                  # ms
a = [0, 0.25, 0.75, 1.0]
                                   # ns
b = 7.0
                   # pA
# Input function
def I_input(t):
   return np.where(t < t1, 0, np.where(t < t2, I0, 0))
# simulation parameters
T = 370
          # ms
dt = 0.01 \# ms
t1 = 20 # ms
t2 = 200  # ms
t = np.arange(0, T, dt)
```

```
# create a grid
     u_{min}, u_{max} = -75, -40
     w_{min}, w_{max} = -45, 70
     N = 15
     \# Create meshgrid for v and w
     u_vals = np.linspace(u_min, u_max, N)
     w_vals = np.linspace(w_min, w_max, N)
     u_grid, w_grid = np.meshgrid(u_vals, w_vals)
     # fromuals (6.3) and (6.4)
     def du_dt(u, w, I, tau_w = 100.0):
         return (-(u-u_rest) + delta_T * np.exp((u - v_rheobase)/delta_T) - R_m * w_
      \hookrightarrow+ R_m * I) / tau_m
     def dw_dt(u, w, a, tau_w = 100.0):
         return (a * (u - u_rest) - w) / tau_w
     # nullcline functions (set (6.3) & (6.4) to zero and solve for w)
     def u_nullcline(u, w, I, tau_w = 100.0):
         return I + 1/R_m*delta_T * np.exp((u - v_rheobase)/delta_T) - 1/R_m*(u -
      ⊶u rest)
     def w_nullcline(u, w, a, tau_w = 100.0):
         return a*(u-u_rest)
```

1. How do the nullclines change with respect to a?

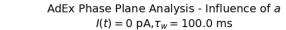
```
[23]: # get the nullclines and vectorfield
w_nc_u = u_nullcline(u_vals, w_vals, 0.0)
w_nc_w_vals = [w_nullcline(u_vals, w_vals, a_i) for a_i in a]
du = du_dt(u_grid, w_grid, 0)
dw_vals = [dw_dt(u_grid, w_grid, a_i) for a_i in a]
```

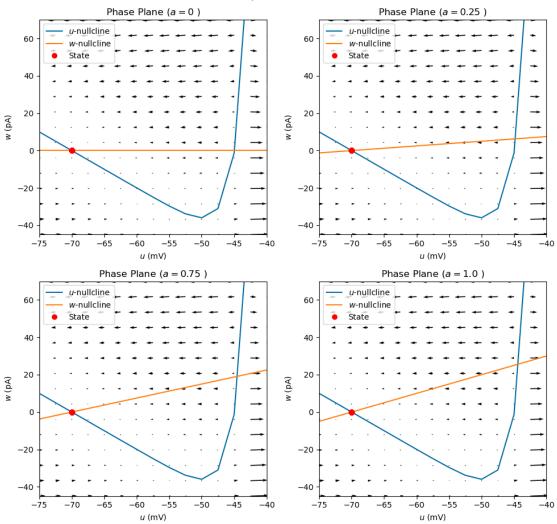
```
[25]: # plot the results
      fig, axs = plt.subplots(2, len(a)//2, figsize = (5*len(a)/2, 5*2))
      plt.suptitle('AdEx Phase Plane Analysis - Influence of $a$ \n$I(t)=0$ pA,' +11
      →r'$\tau_w= {}$ ms'.format(tau_w), fontsize = 14)
      for ax, dw_val, w_nc_w_val, a_i, state in zip(axs.flatten(), dw_vals, u
       →w_nc_w_vals, a, states):
          # plot the vectorfield
         ax.quiver(u_grid, w_grid, du, dw_val)
         # plot the nullclines
         ax.plot(u_vals, w_nc_u, label = '$u$-nullcline')
         ax.plot(u_vals, w_nc_w_val, label = '$w$-nullcline')
          # plot the state
         state_monitor, spike_monitor = state
         ax.plot(state_monitor.v[0]*1e3, state_monitor.w[0]*1e12, 'ro', label = ___
       # lay-out the axes
         ax.set_xlim((u_min, u_max))
         ax.set ylim((w min, w max))
```

```
ax.set_title('Phase Plane ($a = {}$ )'.format(a_i))
ax.set_xlabel('$u$ (mV)')
ax.set_ylabel('$w$ (pA)')

ax.legend()

plt.tight_layout()
plt.show()
```





2. How do the nullclines change if a constant current I(t) = c is applied?

```
[26]: # input currents
              c = [30, 42.0, 45.0, 60.0] # pA
              # get the nullclines and vectorfield
              w_nc_u_vals = [u_nullcline(u_vals, w_vals, c_i) for c_i in c]
              w_nc_w = w_nullcline(u_vals, w_vals, a[1])
              du_vals = [du_dt(u_grid, w_grid, c_i) for c_i in c]
              dw = dw_dt(u_grid, w_grid, a[1])
[27]: # simulate the model
              t_start, t_end = 20, T + 1
              A = 0.0
              I_inputs = [input_factory.get_step_current(t_start, 500, b2.ms, c_i*b2.pA) for_u
                ⇔c_i in c]
              states = [AdEx.simulate_AdEx_neuron(I_stim = I, simulation_time = T * b2.ms,
                                                                                                                                                            v reset = u reset * b2.
                →mV, v_rest = u_rest * b2.mV, v_rheobase = v_rheobase * b2.mV, delta_T = u_rest * b2.mV, delta_

delta_T * b2.mV,
                                                                                                                                                           tau_m = tau_m * b2.ms,__
                 \rightarrowtau_w = tau_w * b2.ms,
                                                                                                                                                           a = a[1] * b2.nS, b = 
                 47 * b2.pA,
                                                                                                                                                           R = R_m * b2.Gohm) for_{\sqcup}
                  →I in I_inputs]
[28]: colors = ['#2ca02c', '#d62728', '#9467bd',
                                       '#8c564b', '#e377c2', '#7f7f7f', '#bcbd22', '#17becf',
                                        '#aec7e8', '#ffbb78', '#98df8a', '#ff9896', '#c5b0d5',
                                        '#c49c94', '#f7b6d2', '#c7c7c7', '#dbdb8d', '#9edae5']
              def plot_vectorfield(ax, grid_vals, u_v_vals, du_dw_vals, ncs, u_w_min_max = u
                 _{\rightarrow}(0, 0, 0, 0), legend = False):
                        w_nc_u, w_nc_w = ncs
                        du, dw = du_dw_vals
                        u_grid, w_grid = grid_vals
                        u_vals, w_vals = u_v_vals
                        u_min, u_max, w_min, w_max = u_w_min_max
                        # plot the vectorfield
                        ax.quiver(u_grid, w_grid, du, dw)
                        # plot the nullclines
                        ax.plot(u_vals, w_nc_u, label = '$u$-nullcline')
                        ax.plot(u_vals, w_nc_w, label = '$w$-nullcline')
```

```
if legend:
        ax.legend()
def plot_state(ax, state):
    state_monitor, spike_monitor = state
    u, w = state_monitor.v[0]*1e3, state_monitor.w[0]*1e12
    t = np.array(state_monitor.t) * 1e5
    t_spikes = np.array(spike_monitor.t) * 1e5
    for i in np.arange(len(t_spikes)):
        if i == 0:
             condition = t < t_spikes[i]</pre>
             ax.plot(u[condition], w[condition], color = colors[i])
        elif i == len(t_spikes):
             condition = t > t_spikes[i]
             ax.plot(u[condition], w[condition], color = colors[i])
        else:
            condition = (t_spikes[i-1] < t) * (t < t_spikes[i])</pre>
            ax.plot(u[condition], w[condition], color = colors[i])
    if len(t_spikes) == 0:
        ax.plot(u[-1], w[-1], 'ro')
def plot_spikes(axs, state):
    ax1, ax2 = axs
    state_monitor, spike_monitor = state
    t = np.array(state_monitor.t) * 1e3
    t_spikes = np.array(spike_monitor.t) * 1e3
    u, w = state_monitor.v[0] * 1e3, state_monitor.w[0] * 1e12
    for i in np.arange(len(t_spikes)):
        if i == 0:
             condition = t <= t_spikes[i]</pre>
             ax1.plot(t[condition], u[condition], color = colors[i])
             ax2.plot(t[condition], w[condition], color = colors[i])
        elif i == len(t_spikes):
             condition = t >= t_spikes[i]
             ax1.plot(t[condition], u[condition], color = colors[i])
             ax2.plot(t[condition], w[condition], color = colors[i])
```

```
[29]: # Create a list to store references to the figures
     figures = []
     for i in range(4):
         # Create a new figure for each block
         fig = plt.figure(figsize = (12, 6))
         figures.append(fig) # Store reference to the figure
         # Add title for the block
         plt.suptitle(f'AdEx Analyisis for $I(t)={c[i]:.2f}$ pA', fontsize = 14)
         # add axes to the figure
         ax11 = fig.add_subplot(2, 2, 1)
         ax12 = fig.add_subplot(2, 2, 3, sharex = ax11)
         ax2 = fig.add_subplot(1, 2, 2)
         state = states[i]
         du = du_vals[i]
         w_nc_u = w_nc_u_vals[i]
         c_i = c[i]
         # plot the vectorfield
         plot_vectorfield(ax2, (u_grid, w_grid), (u_vals, w_vals), (du, dw),__
       plot state(ax2, state)
         plot_spikes((ax11, ax12), state)
```

```
# lay-out the axes
ax2.set_xlim((u_min, u_max))
ax2.set_ylim((w_min, w_max))

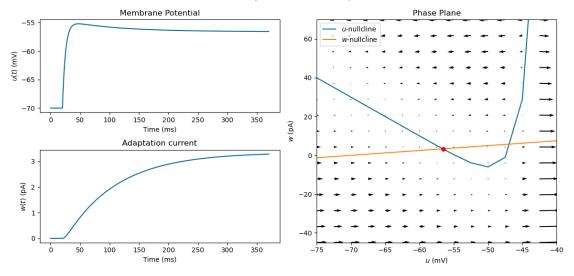
ax2.set_title('Phase Plane')
ax2.set_xlabel('$u$ (mV)')
ax2.set_ylabel('$w$ (pA)')

fig.tight_layout()

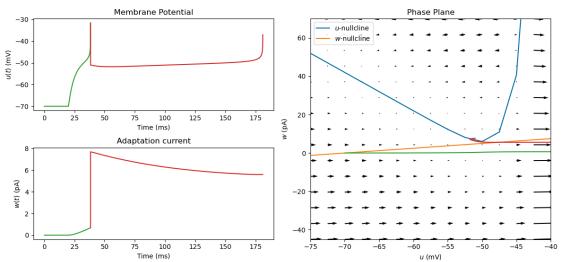
# Adjust space between subplots
#plt.subplots_adjust(hspace=0.5)

# Plot all figures below each other
for fig in figures:
    plt.show(fig)
```

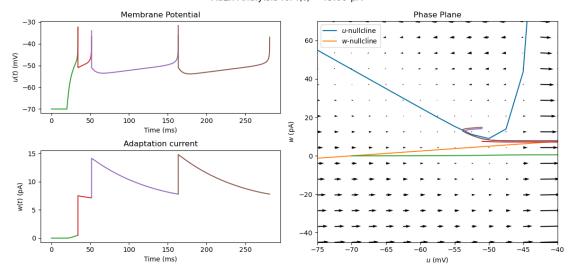
AdEx Analysis for I(t) = 30.00 pA



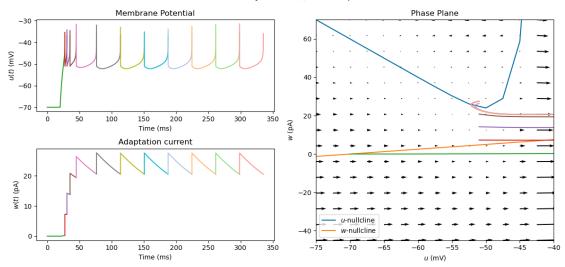
AdEx Analysiis for I(t) = 42.00 pA



AdEx Analysiis for I(t) = 45.00 pA



AdEx Analysiis for I(t) = 60.00 pA



```
[]: # THIS CELL GENERATES A VIDEO, IT TAKES A WHILE TO RENDER THE VIDEO #
     from matplotlib.animation import FuncAnimation, FFMpegWriter
     # Parameters for the simulation
     state = states[-1]
     state_monitor, spike_monitor = state
     t = np.array(state_monitor.t) * 1e3
     t_spikes = np.array(spike_monitor.t) * 1e3
     u, w =state_monitor.v[0] * 1e3, state_monitor.w[0] * 1e12
     w_min2, w_max2 = float(min(w)), float(max(w))
     u_min2, u_max2 = float(min(u)), float(max(u))
     du = du_vals[-1]
     w_nc_u = w_nc_u_vals[-1]
     c_i = c[-1]
     # Create a new figure for each block
     fig = plt.figure(figsize = (12, 6))
     # Add title for the block
     plt.suptitle(f'AdEx Analyisis for $I(t)={c[i]:.2f}$ pA', fontsize = 14)
     # add axes to the figure
     ax11 = fig.add_subplot(2, 2, 1)
     ax12 = fig.add_subplot(2, 2, 3)
     ax2 = fig.add_subplot(1, 2, 2)
```

```
plot_vectorfield(ax2, (u_grid, w_grid), (u_vals, w_vals), (du, dw), (w_nc_u,_
 \hookrightarrow w_nc_w))
ax11.set title('Membrane Potential')
ax12.set_title('Adaptation Current')
ax11.set_xlabel('Time (ms)')
ax12.set_xlabel('Time (ms)')
ax11.set_ylim((u_min2, u_max2))
ax12.set_ylim((w_min2 , w_max2))
ax11.set_xlim((0, max(t)))
ax12.set_xlim((0, max(t)))
ax2.set_title('Phase Plane')
ax2.set_xlabel('$u$ (mV)')
ax2.set_ylabel('$w$ (pA)')
ax2.set_xlim((u_min , u_max))
ax2.set_ylim((w_min, w_max + 5))
fig.tight_layout()
def update(frame):
    frame = int(frame)
    ax2.plot(u[:frame], w[:frame], color = 'red')
    ax11.plot(t[:frame], u[:frame], color = 'red')
    ax12.plot(t[:frame], w[:frame], color = 'red')
    return None
ani = FuncAnimation(fig, update, frames = np.arange(0, len(t) + 100, 100),
→interval = 100)
writer = FFMpegWriter(fps = 10, metadata = dict(artist = 'Constantijnu

    Goppers'), bitrate = 1800)
ani.save('AdeX_analyis_.mp4', writer = writer)
```

3. Interpretation of b (pA)

```
[30]: # b-values
b_vals = [3.0, 7.0, 10.0, 25.0]
```

```
# get the nullclines and vectorfield
w_nc_u = u_nullcline(u_vals, w_vals, c[-1])
w_nc_w = w_nullcline(u_vals, w_vals, a[1])
du = du_dt(u_grid, w_grid, c[-1])
dw = dw_dt(u_grid, w_grid, a[1])
# simulate the model
#t start, t end = 20, T + 1
\#I\_inputs = [input\_factory.get\_step\_current(t\_start, 500, b2.ms, c\_i*b2.pA) for_{\sqcup} 
    \hookrightarrow c i in c]
states = [AdEx.simulate_AdEx_neuron(I_stim = I_inputs[-1], simulation_time = T_
      →* b2.ms,
                                                                                                                                                                                                                                                                                                                                        v_reset = u_reset * b2.
      ⇒mV, v_rest = u_rest * b2.mV, v_rheobase = v_rheobase * b2.mV, delta_T = ∪

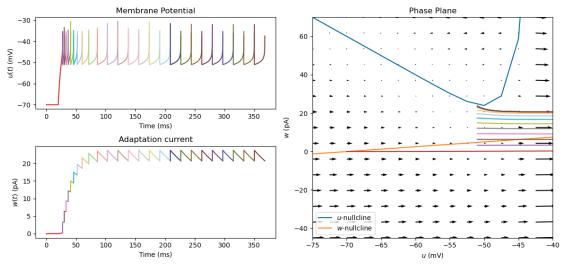
delta_T * b2.mV,
                                                                                                                                                                                                                                                                                                                                       tau_m = tau_m * b2.ms,_{\sqcup}

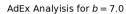
ytau_w = tau_w * b2.ms,
                                                                                                                                                                                                                                                                                                                                       a = a[1] * b2.nS, b = 
      \rightarrow b * b2.pA,
                                                                                                                                                                                                                                                                                                                                       R = R_m * b2.Gohm) for_{\sqcup}
       →b in b_vals]
colors = ['#d62728', '#9467bd',
```

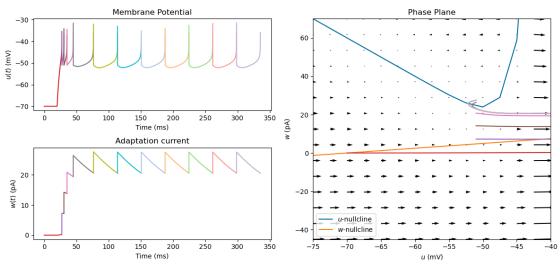
```
[31]: # Create a list to store references to the figures
                '#8c564b', '#e377c2', '#7f7f7f', '#bcbd22', '#17becf',
                '#aec7e8', '#ffbb78', '#98df8a', '#ff9896', '#c5b0d5',
                '#c49c94', '#f7b6d2', '#c7c7c7', '#dbdb8d', '#9edae5',
                '#393b79', '#637939', '#8c6d31', '#843c39', '#7b4173',
                '#5254a3', '#637939', '#8c6d31', '#843c39', '#7b4173',
                '#393b79', '#5254a3', '#6b6ecf', '#9c9ede', '#637939',
                '#8c6d31', '#843c39', '#7b4173', '#bd9e39', '#d6616b',
                '#ce6dbd', '#9c9ede', '#edc948', '#8ca252', '#b5cf6b',
                '#c49c94', '#e7ba52', '#e7969c', '#d6616b', '#7b4173']
      figures = []
      for i in range(len(b_vals)):
          # Create a new figure for each block
          fig = plt.figure(figsize = (12, 6))
          figures.append(fig)
          # Add title for the block
          plt.suptitle(f'AdEx Analysis for $b={b_vals[i]}$', fontsize = 14)
```

```
# add axes to the figure
   ax11 = fig.add_subplot(2, 2, 1)
   ax12 = fig.add_subplot(2, 2, 3)
   ax2 = fig.add_subplot(1, 2, 2)
   state = states[i]
   # plot the vectorfield/...
   plot_vectorfield(ax2, (u_grid, w_grid), (u_vals, w_vals), (du, dw),_u
 plot_state(ax2, state)
   plot_spikes((ax11, ax12), state)
   # lay-out the axes
   ax2.set_xlim((u_min, u_max))
   ax2.set_ylim((w_min, w_max))
   ax2.set_title('Phase Plane')
   ax2.set_xlabel('$u$ (mV)')
   ax2.set_ylabel('$w$ (pA)')
   fig.tight_layout()
# Plot all figures below each other
for fig in figures:
   plt.show(fig)
```

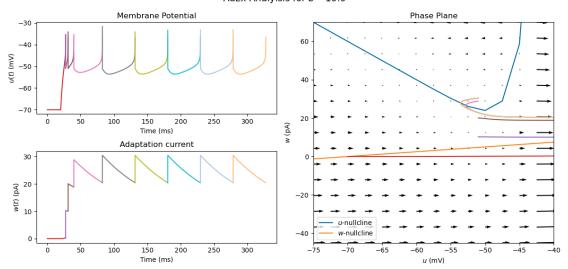




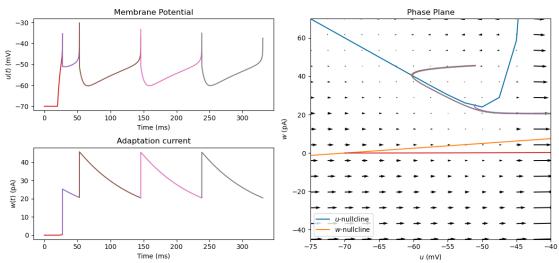




AdEx Analyisis for b = 10.0



AdEx Analyisis for b = 25.0



4. Influence of τ_w

```
[32]: # tau values
      tau_vals = [10.0, 50.0, 200.0] # ms
      # get the nullclines and vectorfield
      w_nc_u_vals = [u_nullcline(u_vals, w_vals, c[-1], tau_w = tau) for tau in_
       →tau_vals]
      w_nc_w_vals = [w_nullcline(u_vals, w_vals, a[1], tau_w = tau) for tau in_
       →tau_vals]
      du_vals = [du_dt(u_grid, w_grid, c[-1], tau_w = tau) for tau in tau_vals]
      dw_vals = [dw_dt(u_grid, w_grid, a[1], tau_w = tau) for tau in tau_vals]
      # simulate the model
      \#t_start, t_end = 20, T + 1
      #I inputs = [input factory.get_step_current(t_start, 500, b2.ms, c_i*b2.pA) for_
      states = [AdEx.simulate_AdEx_neuron(I_stim = I_inputs[-1], simulation_time = T_i
       \rightarrow* b2.ms,
                                                                  v_reset = u_reset * b2.
       omV, v_rest = u_rest * b2.mV, v_rheobase = v_rheobase * b2.mV, delta_T = ∪

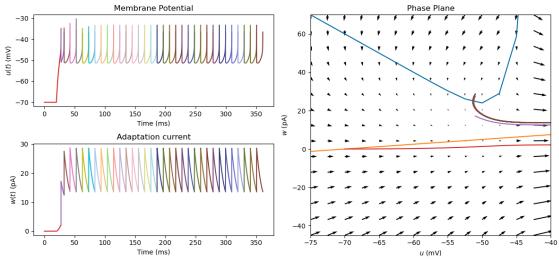
delta_T * b2.mV,
                                                                  tau_m = tau_m * b2.ms,_
       \rightarrowtau_w = tau * b2.ms,
                                                                  a = a[1] * b2.nS, b = 
       \hookrightarrow15 * b2.pA,
```

```
R = R_m * b2.Gohm) for_{\sqcup} \hookrightarrowtau in tau_vals]
```

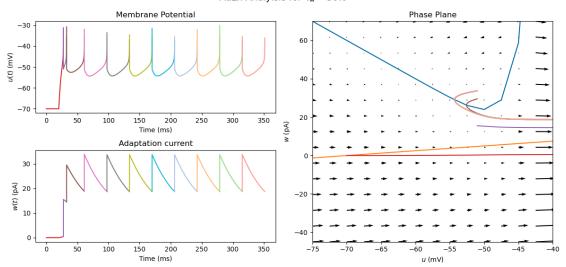
```
[33]: # Create a list to store references to the figures
      figures = []
      for i in range(len(tau_vals)):
          # Create a new figure for each block
          fig = plt.figure(figsize = (12, 6))
          figures.append(fig)
          # Add title for the block
          plt.suptitle(f'AdEx Analyisis for ' + r'$\tau_w='+f'{tau_vals[i]}$',__
       ⇒fontsize = 14)
          # add axes to the figure
          ax11 = fig.add_subplot(2, 2, 1)
          ax12 = fig.add_subplot(2, 2, 3)
          ax2 = fig.add_subplot(1, 2, 2)
          state = states[i]
          du, dw = du_vals[i], dw_vals[i]
          w_nc_u, w_nc_w = w_nc_u_vals[i], w_nc_w_vals[i]
          c_i = c[-1]
          # plot the vectorfield/...
          plot_vectorfield(ax2, (u_grid, w_grid), (u_vals, w_vals), (du, dw),__

    (w_nc_u, w_nc_w))
          plot_state(ax2, state)
          plot_spikes((ax11, ax12), state)
          # lay-out the axes
          ax2.set_xlim((u_min, u_max))
          ax2.set_ylim((w_min, w_max))
          ax2.set_title('Phase Plane')
          ax2.set_xlabel('$u$ (mV)')
          ax2.set_ylabel('$w$ (pA)')
          fig.tight_layout()
      # Plot all figures below each other
      for fig in figures:
          plt.show(fig)
```

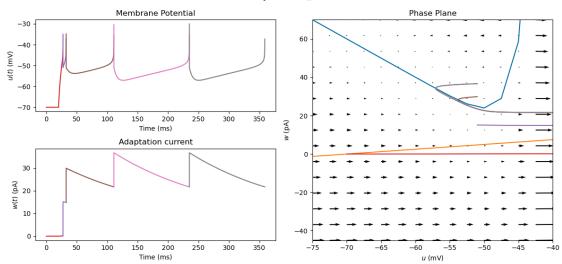




AdEx Analyisis for $\tau_{w} = 50.0$



AdEx Analysiis for $\tau_w = 200.0$



4.1 Answer:

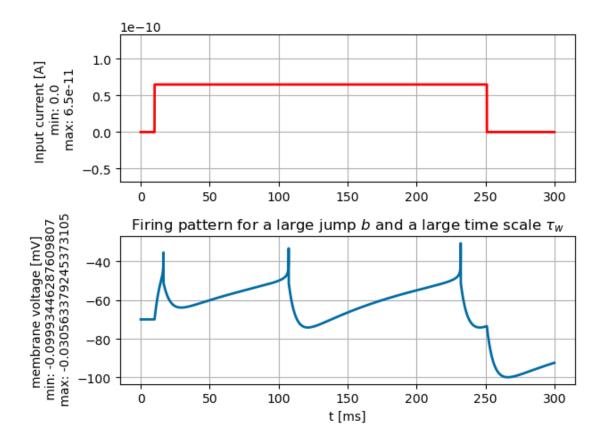
- 1. How do the nullclines change with respect to a?

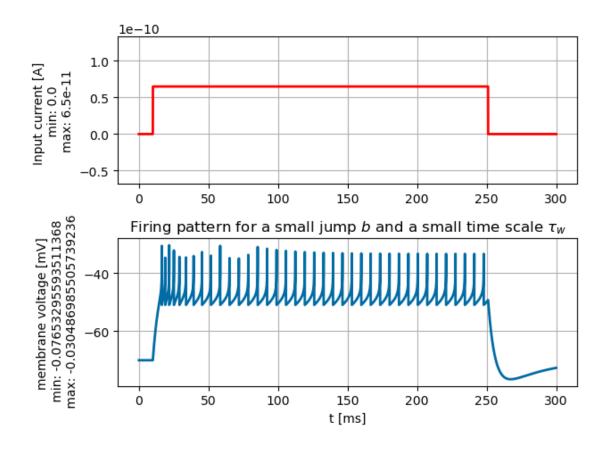
 The parameter a is the slope of the linear w-nullcline. Increasing |a| increases the slope of the w-nullcline. A does not affect the u-nullcline.
- 2. How do the nullclines change if a constant current I(t) = c > 0 is applied? The *u*-nullcline shifts upwards (downwards) for increasing (decraesing) input current.
- 3. What is the interpretation of parameter b?

 The parameter b is the spike triggered current that depolarizes (b < 0) or polarizes (b > 0) the neuron. In the phase plane it corresponds to a vertical jump in the state. The jump is proportional to |b|.
- 4. How do flow arrows change as τ_w gets bigger? The flow arrows are more oriented along the u axis (horizontal).

3.4.3 A4.2 Predict firing pattern

• Go back to Q4.2





4.2 Answer:

The a parameter is the coupeling between the adaptation current w and the membrane potential u. A value for a of about 0.01 nS corresponds to an almost horizontal (linear) w-nullciline. I.e. there is almost no coupeling of w with u.

A large jump b will result in a larger time interval between spikes and thus the firing rate will be lower. If b is lower it takes more steps upwards in the phase plane to come in a space where the vertical vectors are more dominant and cause a small detour in the phase plane 1 . As stated in 4.1, a larger τ_w value causes more horizonatal oriented vectors and therefore we expect that for large τ_w values the detour of the state in the phase plane will be higher, which will correspond with a larger interspike interval.

These effects are visible in the figure above: for small b and large τ_w there are less spikes generated.

¹ see the video (AdEx_analysis.mp4) I have made and look at how the state curve differs (in time) from bottom to top in the phase plane. You van see that around u-nullcline, the vectors are smaller in magnitude, so as the state makes jumps (of amount b), it migrates to a space where it is less pulled to the right and thus it will take more time to generate a spike.