

Total
10 points

Problem Set 1 Solution

1. Membrane model.

When time-varying current is injected into a cell, the equilibrium voltage is governed by this equation:

1 point

$$C_m \frac{dV}{dt} + (V - V_{leak})G_{leak} - I_{in}(t) = 0$$

Solving this differential equation, voltage can be expressed as a function of the time-varying current via convolution:

$$V = V_{leak} + \left\{ I_{in} * \frac{1}{C_m} e^{-t/\tau} \right\}$$

(a) Uniform current.

1 point

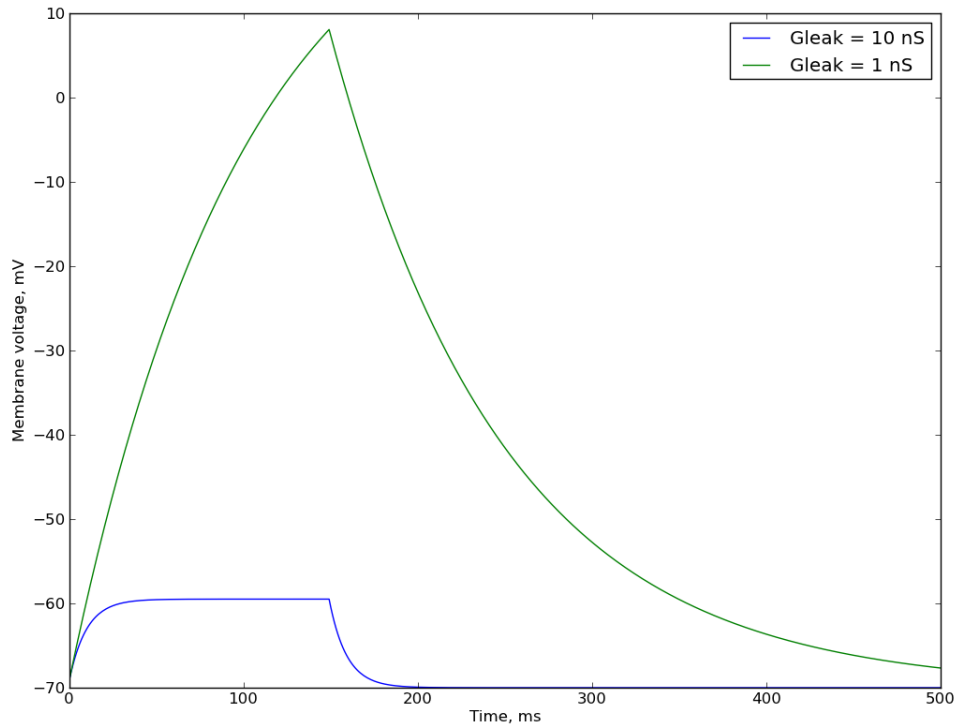


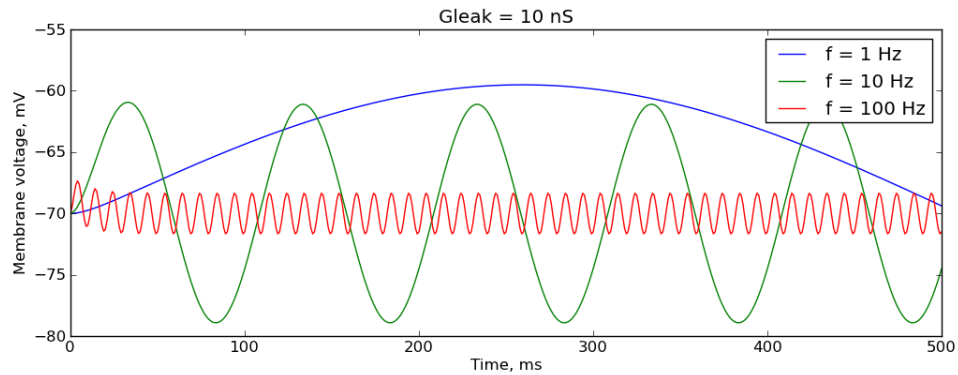
Figure 1: Problem 1(a).

0.5 point

Injecting a constant current for 150 ms causes the membrane voltage to increase from the leak potential of -70mV . For $G_{leak} = 10\text{nS}$, the voltage saturates at -60mV and falls off after the current is turned off. For $G_{leak} = 1\text{nS}$, the voltage does not reach saturation before the current is turned off. It peaks at approximately 8mV and decreases after the current is turned off.

(b) Sinewave current.

1 point



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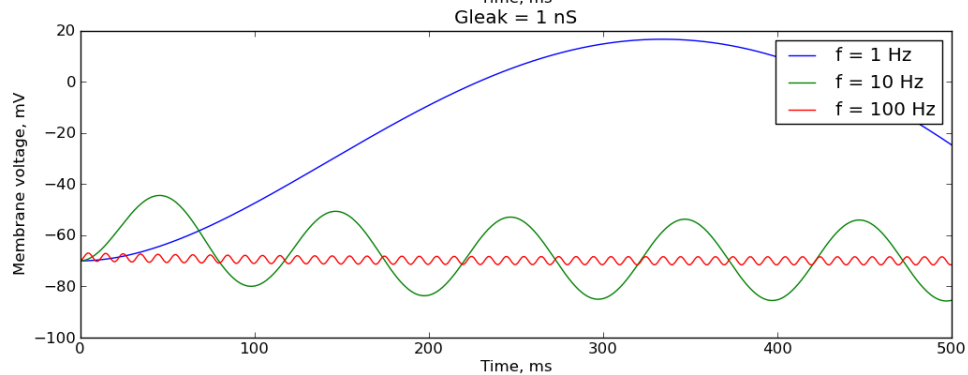


Figure 2: Problem 1(b).

0.5 point

Injecting a sinusoidal current produces a sinusoidal change in voltage. The frequency of the voltage fluctuation is equal to the frequency of the current. The amplitude and phase are dependent on the conductance: the amplitude is smaller for the higher conductance, but the phase shift is higher for the smaller conductance.

(c) White Gaussian noise.

1 point

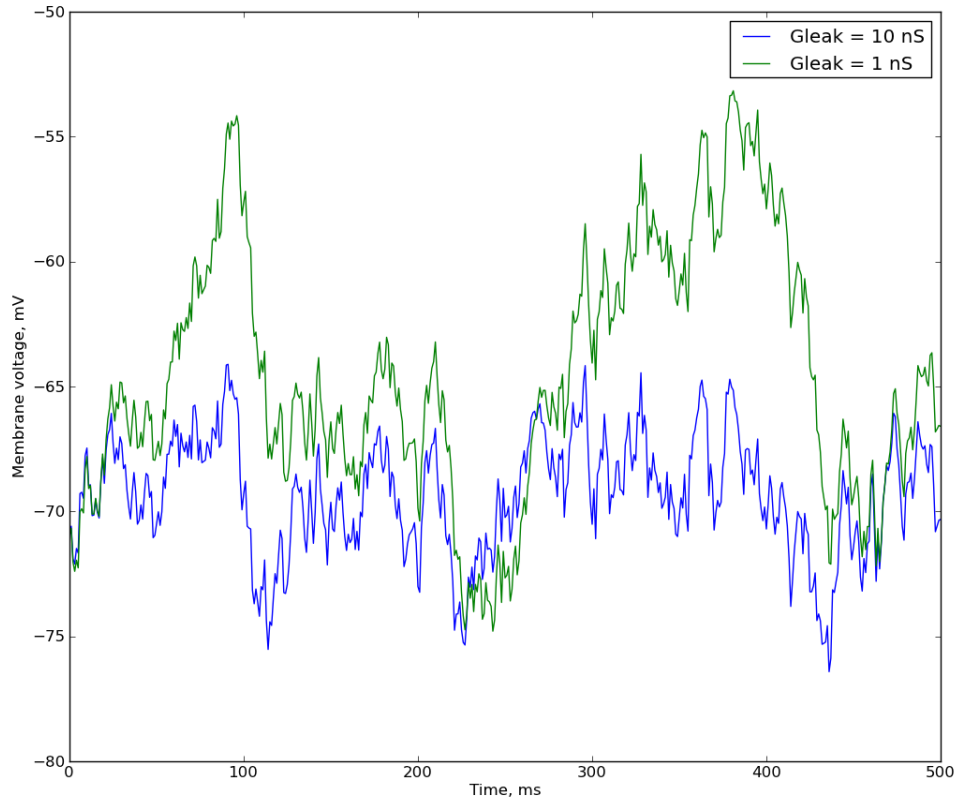


Figure 3: Problem 1(c).

Injecting a white noise current produces greater fluctuations in voltage for the lower conductance.

2. Shunting inhibition.

At equilibrium, the membrane voltage can be calculated from the reversal potentials and conductances of the ion channels. Taking into account only the sodium and chloride channels and leak voltage and conductance, the equation for calculating the membrane potential is:

$$V = \frac{V_{leak}G_{leak} + V_{Na}G_{Na} + V_{Cl}G_{Cl}}{G_{leak} + G_{Na} + G_{Cl}}$$

1 point

2 points

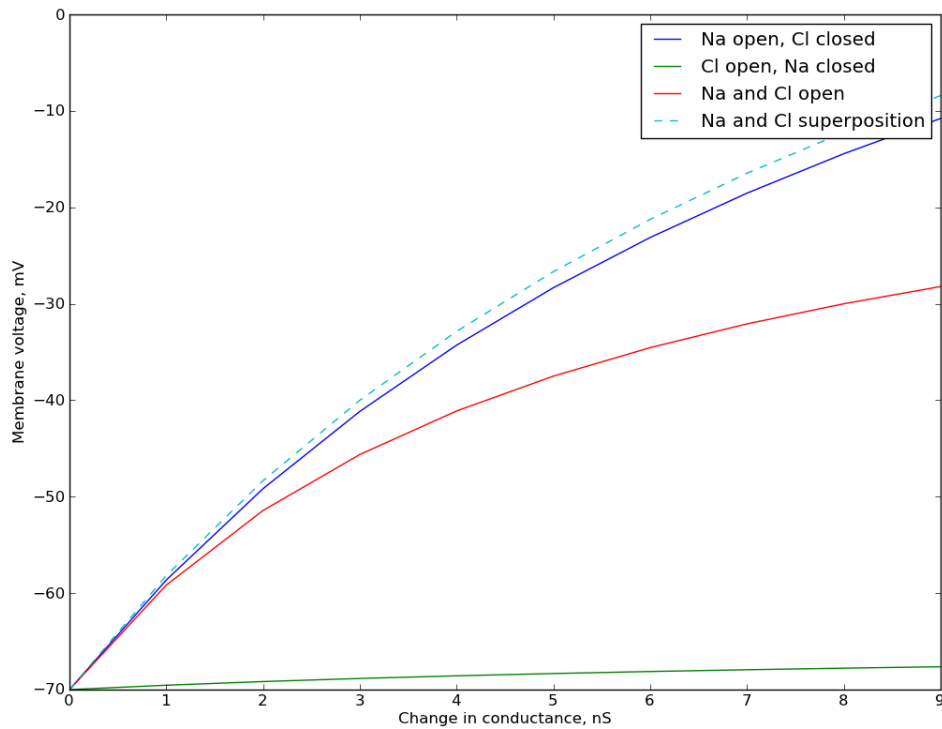
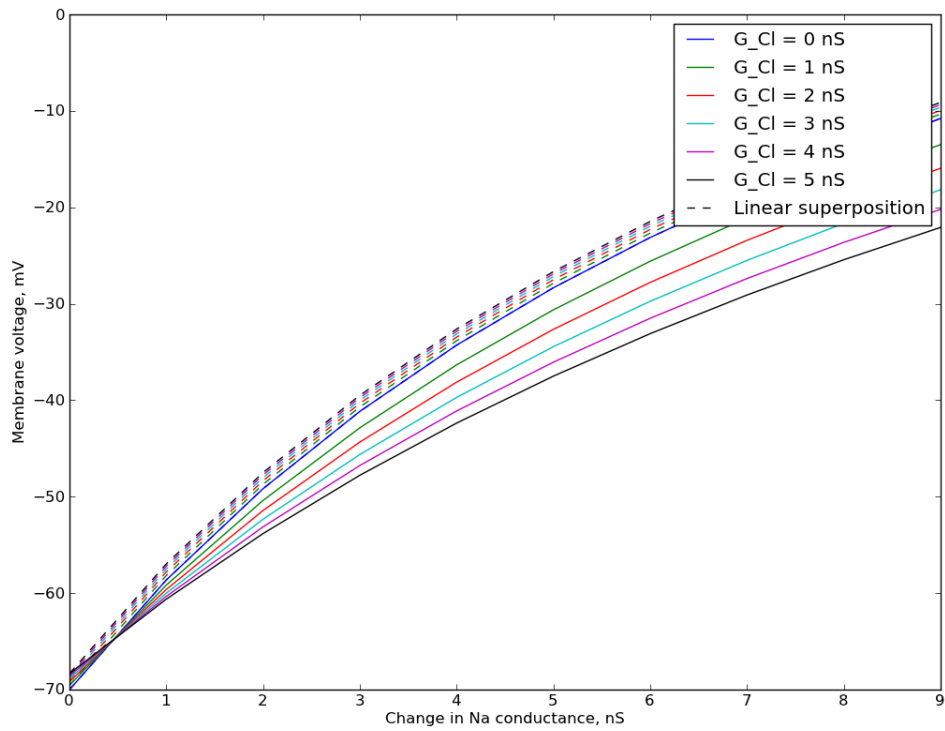


Figure 4: Problem 2.

1 point

Opening Na channels and keeping Cl channels closed produces a fairly large positive change in voltage, whereas opening only Cl channels and keeping Na channels closed produces a much smaller positive change. Opening both Na and Cl channels produces a smaller positive change than opening Na channels only. Linear superposition predicts a much higher change in voltage, suggesting that the interaction of Na and Cl channel opening is nonlinear.



Gold Star

Figure 5: Problem 2.

Greater Cl conductance inhibits the positive effect of increasing Na conductance on the membrane voltage, whereas linear superposition predicts the opposite effect. This result further highlights the nonlinear interaction between Na and Cl conductance.

Other students worked with: Alina Liberman, Anwar Nunez-Elizalde

Code:

```
1 import numpy as np
2 from matplotlib.pyplot import figure, savefig
3
4 ## Problem 1.
5 ## Given:
6 Vleak = -70 # leak voltage, mV
7 Cm = 100.e-12 # membrane capacitance, pF
8 Gleak1 = 10.e-9 # leak conductance 1, nS
9 Gleak2 = 1.e-9 # leak conductance 2, nS
10 tau1 = Cm/Gleak1 # time constant for conductance 1
11 tau2 = Cm/Gleak2 # time constant for conductance 2
12 nT = 501 # for 500 ms
13 tps = np.array(range(nT))*1.e-3 # the timepoints
14
15 ## Voltage for two different conductance values.
16 V1 = np.zeros((nT,))
17 V2 = np.zeros((nT,))
18
19 ## Initial condition:
20 V1[0] = Vleak
21 V2[0] = Vleak
22
23 ## Problem 1a.
24 ## Input current I0 for 0<=t<150 ms; 0 for t<0, t>150 ms
25 I0 = 100.e-12 # Input current, pA
26 Iin = np.zeros((nT,))
27 Iin[:150] = I0
28
29 ## Voltage is equal to the convolution of injected current and 1/Cm*exp(-t/tau
30 ), plus initial voltage
31 Va1 = Vleak+np.convolve(Iin, 1/Cm*exp(-tps/tau1))[:nT]
32 Va2 = Vleak+np.convolve(Iin, 1/Cm*exp(-tps/tau2))[:nT]
33
34 fa = figure(num=1, figsize=(12, 9), dpi=80, facecolor='w', edgecolor='k')
35 ax = fa.add_subplot(111)
36 va1 = ax.plot(Va1)
37 va2 = ax.plot(Va2)
38 ax.set_ylabel('Membrane voltage, mV')
39 ax.set_xlabel('Time, ms')
40 ax.legend([va1[0], va2[0]], ['Gleak = 10 nS', 'Gleak = 1 nS'])
41 savefig('Problem1a.png')
42
43 ## Problem 1b.
44
45 A = 100.e-12 #Input current, pA
46 f1 = 1 # Hz
47 f2 = 10 # Hz
48 f3 = 100 #Hz
49
50 ## Input current for each frequency:
51 Iin1 = A*sin(2*pi*f1*tps)
```

```

51 Iin2 = A*sin(2*pi*f2*tps)
    Iin3 = A*sin(2*pi*f3*tps)
53
    ## Calculate the current using the same formula as above, for each conductance
    and frequency:
55 Vb_g1_f1 = Vleak+np.convolve(Iin1, 1/Cm*exp(-tps/tau1))[:nT]
    Vb_g1_f2 = Vleak+np.convolve(Iin2, 1/Cm*exp(-tps/tau1))[:nT]
57 Vb_g1_f3 = Vleak+np.convolve(Iin3, 1/Cm*exp(-tps/tau1))[:nT]
59 Vb_g2_f1 = Vleak+np.convolve(Iin1, 1/Cm*exp(-tps/tau2))[:nT]
    Vb_g2_f2 = Vleak+np.convolve(Iin2, 1/Cm*exp(-tps/tau2))[:nT]
61 Vb_g2_f3 = Vleak+np.convolve(Iin3, 1/Cm*exp(-tps/tau2))[:nT]
63 fb = figure(num=2, figsize=(12, 9), dpi=80, facecolor='w', edgecolor='k')
    axb1 = fb.add_subplot(211)
65 axb1.set_title('Gleak = 10 nS')
    vb_g1_f1 = axb1.plot(Vb_g1_f1)
67 vb_g1_f2 = axb1.plot(Vb_g1_f2)
    vb_g1_f3 = axb1.plot(Vb_g1_f3)
69 axb1.set_ylabel('Membrane voltage, mV')
    axb1.set_xlabel('Time, ms')
71 axb1.legend([vb_g1_f1[0], vb_g1_f2[0], vb_g1_f3[0]], ['f = 1 Hz', 'f = 10 Hz',
    'f = 100 Hz'])
73 axb2 = fb.add_subplot(212)
    axb2.set_title('Gleak = 1 nS')
75 vb_g2_f1 = axb2.plot(Vb_g2_f1)
    vb_g2_f2 = axb2.plot(Vb_g2_f2)
77 vb_g2_f3 = axb2.plot(Vb_g2_f3)
    axb2.set_ylabel('Membrane voltage, mV')
79 axb2.set_xlabel('Time, ms')
    axb2.legend([vb_g2_f1[0], vb_g2_f2[0], vb_g2_f3[0]], ['f = 1 Hz', 'f = 10 Hz',
    'f = 100 Hz'])
81 savefig('Problem1b.png')
83 ## Problem 1c.
    ## Injected current is Gaussian noise, st. dev = 100 pA
85 Iinc = 100.e-12*np.random.randn(500,)
87 Vc1 = Vleak + np.convolve(Iinc, 1/Cm*exp(-tps/tau1))[:nT]
    Vc2 = Vleak + np.convolve(Iinc, 1/Cm*exp(-tps/tau2))[:nT]
89
    fc = figure(num=3, figsize=(12, 10), dpi=80, facecolor='w', edgecolor='k')
91 axc = fc.add_subplot(111)
    vc1 = axc.plot(Vc1)
93 vc2 = axc.plot(Vc2)
    axc.set_ylabel('Membrane voltage, mV')
95 axc.set_xlabel('Time, ms')
    axc.legend([vc1[0], vc2[0]], ['Gleak = 10 nS', 'Gleak = 1 nS'])
97 savefig('Problem1c.png')
99 ## Problem 2
    Gleak = 10.e-9 # leak conductance, nS
101 Vleak = -70 # leak voltage, mV

```

```

V_Na = 55 # Na reversal potential, mV
103 V_Cl = -65 # Cl reversal potential, mV

105 ## Equilibrium potential is:
## V = (V_leak*G_leak + V_Na*G_Na + V_Cl*G_Cl)/(G_leak + G_Na + G_Cl)
107
## Changing the sodium conductance alone:
109 G_Na = np.array(range(10))*1.e-9
G_Cl = 0
111 Vn = (Vleak*Gleak + V_Na*G_Na + V_Cl*G_Cl)/(Gleak+G_Na+G_Cl)

113 ## Changing the chloride conductance alone:
G_Na = 0
115 G_Cl = np.array(range(10))*1.e-9
Vc = (Vleak*Gleak + V_Na*G_Na + V_Cl*G_Cl)/(Gleak+G_Na+G_Cl)
117
## Changing both conductances together:
119 G_Na = np.array(range(10))*1.e-9
G_Cl = np.array(range(10))*1.e-9
121 Vnc = (Vleak*Gleak + V_Na*G_Na + V_Cl*G_Cl)/(Gleak+G_Na+G_Cl)

123 f2 = figure(num=4, figsize = (12, 9), dpi = 80, facecolor = 'w', edgecolor = '
k')
ax2 = f2.add_subplot(111)
125 v2n = ax2.plot(Vn)
v2c = ax2.plot(Vc)
127 v2nc = ax2.plot(Vnc)
v2sp = ax2.plot(Vn+Vc-Vleak, '--') # linear superposition, subtracting off
Vleak since it was added twice
129 ax2.legend([v2n[0], v2c[0], v2nc[0], v2sp[0]], ['Na open, Cl closed', 'Cl open
, Na closed', 'Na and Cl open', 'Na and Cl superposition'])
ax2.set_ylabel('Membrane voltage, mV')
131 ax2.set_xlabel('Change in conductance, nS')
savefig('Problem2.png')
133
## Changing GNa for each value of G_Cl between 0 and 5 nS:
135 f2all = figure(num=5, figsize = (12, 9), dpi = 80, facecolor='w', edgecolor='k
')
ax3 = f2all.add_subplot(111)
137 G_Na = np.array(range(10))*1.e-9
leg = []
139 colors = ['b', 'g', 'r', 'c', 'm', 'k']
for ii, G_Cl in enumerate(range(6)):
141 G_Cl = G_Cl*1.e-9
Vnc_i = (Vleak*Gleak + V_Na*G_Na + V_Cl*G_Cl)/(Gleak+G_Na+G_Cl)
143 Vsp_i = (Vleak*Gleak + V_Na*G_Na)/(Gleak+G_Na) + (Vleak*Gleak +V_Cl*G_Cl)/(
Gleak+G_Cl) - Vleak # linear superposition
vnc_i = ax3.plot(Vnc_i, colors[ii])
145 vsp_i = ax3.plot(Vsp_i, colors[ii]+'--')
leg.append(vnc_i[0])
147 leg.append(vsp_i[0])
ax3.set_ylabel('Membrane voltage, mV')
149 ax3.set_xlabel('Change in Na conductance, nS')
ax3.legend(leg, ['G_Cl = 0 nS', 'G_Cl = 1 nS', 'G_Cl = 2 nS', 'G_Cl = 3 nS', '

```



```
151 savefig('Problem2all.png')
    G.Cl = 4 nS', 'G.Cl = 5 nS', 'Linear superposition']])
```

hw1.py