

Homework #3.**Problem 1a.**

```

1  %Define constants;
2  R=8.314;
3  T=37+273.15;
4  F=96485;
5  z_K=1;
6  z_Ca=2;
7  z_Cl=-1;
8  z_Na=1;
9  %Define concentrations external and internal
10 K_ext=5;
11 K_int=100;
12 Na_ext=150;
13 Na_int=15;
14 Ca_ext=2;
15 Ca_int=0.0002;
16 Cl_ext=150;
17 Cl_int=13;
18 %compute the reversal potential for each ion
19 E_K=(R*T/(z_K*F))*log(K_ext/K_int);
20 E_Na=(R*T/(z_Na*F))*log(Na_ext/Na_int);
21 E_Ca=(R*T/(z_Ca*F))*log(Ca_ext/Ca_int);
22 E_Cl=(R*T/(z_Cl*F))*log(Cl_ext/Cl_int);
23
24 %-----RESULTS:-----
25 %          E_K =  -0.0801 V
26 %          E_Na =   0.0615 V
27 %          E_Ca =   0.1231 V
28 %          E_Cl =  -0.0654 V
29 %-----RESULTS:-----
30

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Problem 1b.

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31 - syms Rm pK pNa K_in K_ext Na_ext Na_in R T F
32 - pk=solve(Rm == R*T/F*log((pK*K_ext +pNa*Na_ext)/(pK*K_in +pNa*Na_in)),pK);
33
34 %Define constants;
35 R=8.314;
36 T=37+273.15;
37 F=96485;
38 Rm= -0.065;
39
40 %Define concentrations external and internal
41 K_ext=5;
42 K_int=100;
43 Na_ext=150;
44 Na_int=15;
45 %the formula for pK is as follows in terms of variables and constants...
46 -(Na_ext*pNa - Na_int*pNa*exp((F*Rm)/(R*T)))/(K_ext - K_int*exp((F*Rm)/(R*T)))
47 %-----RESULTS:-----
48 % as a function of sodium permeability and concentrations of sodium and
49 % potassium, the formula for the permeability of K is as follows
50 % pk = (892807347126063155*pNa)/22726524804436992
51 % pk = 39.2848*pNa
52 %
53 %-----RESULTS:-----

```

Problem 2:

$$\frac{dn}{dt} + \beta_n n = \alpha_n (1-n)$$

$$\frac{dn}{dt} + (\alpha_n + \beta_n)n = \alpha_n$$

$$a_1 = 1 \quad a_0 = \alpha_n + \beta_n$$

$$1 \cdot \frac{d}{dt} K e^{st} + (\alpha_n + \beta_n) K e^{st} = 0$$

$$K s e^{st} + (\alpha_n + \beta_n) K e^{st} = 0 \rightarrow s = -\alpha_n - \beta_n$$

$$s + (\alpha_n + \beta_n) = 0 \rightarrow s = -(\alpha_n + \beta_n)$$

$$n_i = K e^{-t/\tau}$$

$$\frac{dn}{dt} + (\beta_n + \alpha_n)n = \alpha_n = 0$$

$$K e^{0t} + (\beta_n + \alpha_n)n_s = \alpha_n = 0$$

$$n_s = \frac{\alpha_n}{\alpha_n + \beta_n}$$

$$n = n_i + n_s = K e^{-t/\tau} + \frac{\alpha_n}{\alpha_n + \beta_n}$$

Initial Cond

$$n = n_0 \text{ @ } t=0$$

$$\text{@ } t=0 \quad n(t=0) = K + \frac{\alpha_n}{\alpha_n + \beta_n} = n_0$$

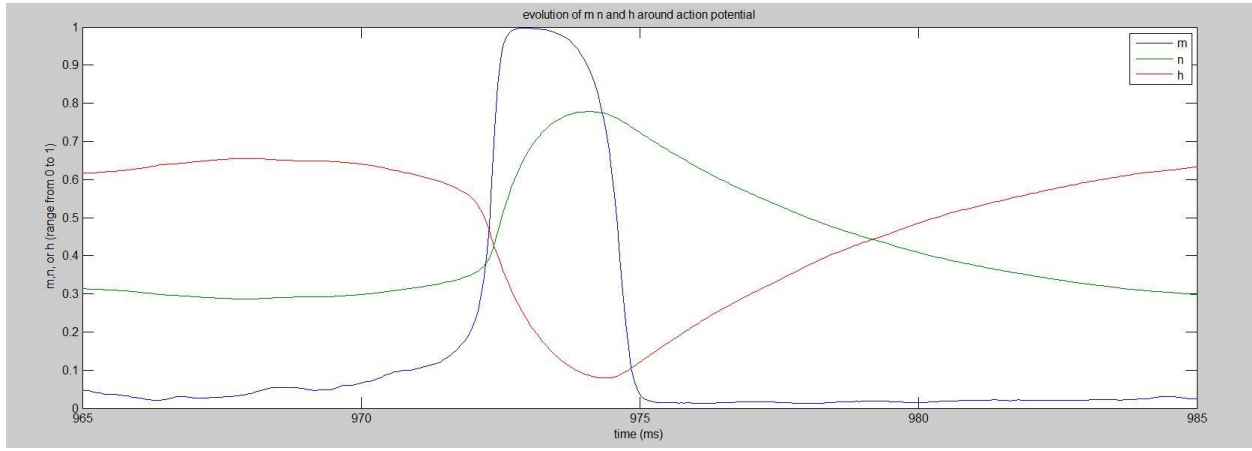
$$K = n_0 - \frac{\alpha_n}{\alpha_n + \beta_n} = n_0 - n_\infty$$

$$n = (n_0 - n_\infty) e^{-t/\tau} + n_\infty$$

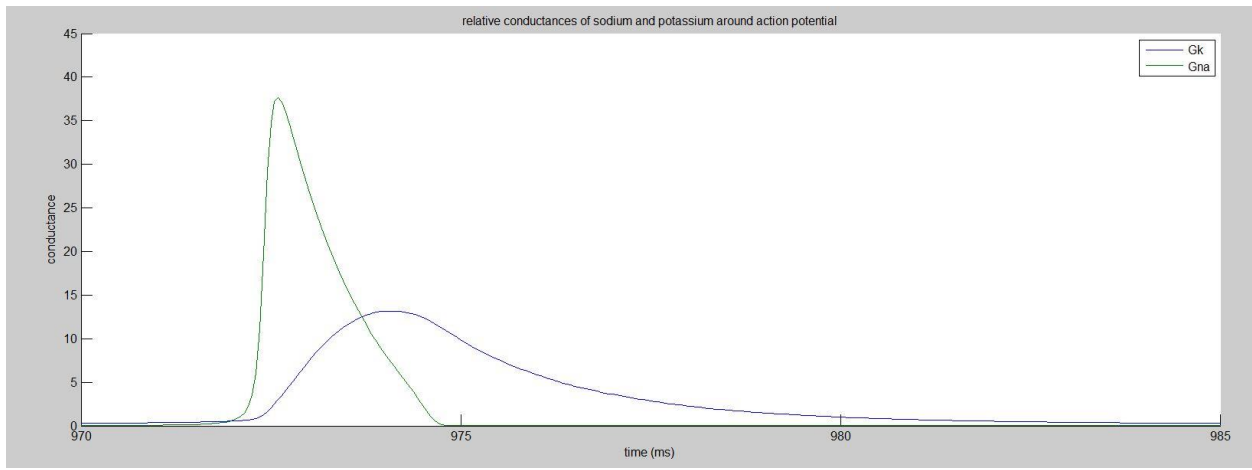


Problem 3:

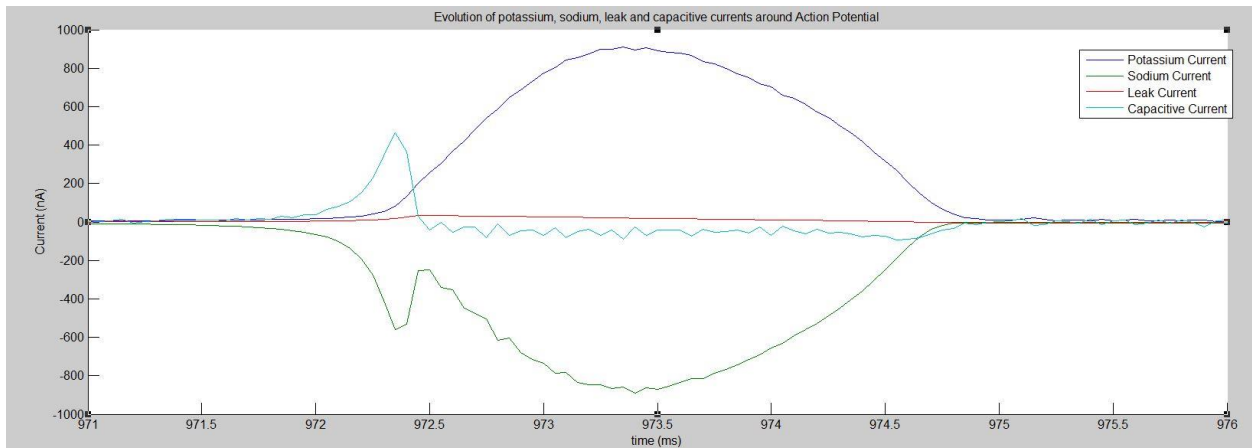
(i):



(ii):



(iii):

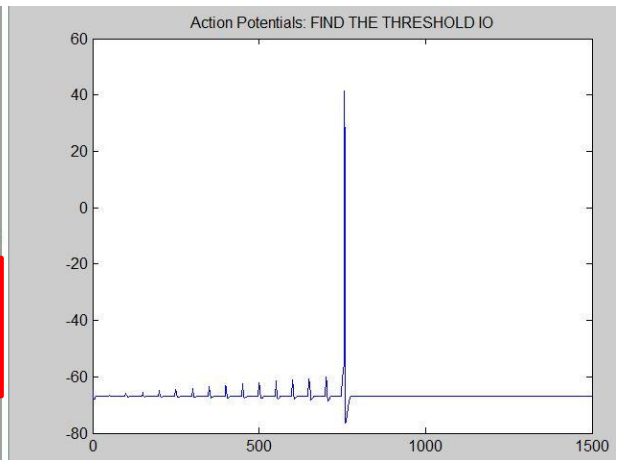


Threshold was determined by increasing the value of I_0 at visible spacings on the graph until an AP presented itself, then narrowing the range down back and forth until a precise value was obtained. Here the value of threshold is $I_0 = 9.758 * 15 = 146.7705 \text{ nA}$

```

4 - clear
5 - c1c
6 - T = 1500; dt = 0.05;
7 - Tsteps = round(T/dt);
8 - C = 1; % capacitance (microF/cm^2)
9 - V1 = -77; % leak reversal potential (mV)
10 - g1 = 0.3; % leak conductance (mS/cm^2)
11 - VNa = 55; % Na reversal potential
12 - gNa = 120; % open Na conductance
13 - VK = -77; % K reversal potential
14 - V1 = -61; % leak reversal potential (mV)
15 - gK = 36; % open K conductance
16 - % input current: sets length of applied current to 0 and randomizing
17 - % current amplitude across 0-10 for 4 and 5 comment out the 2nd line and
18 - % redefine current with a vector of currents to find the threshold current.
19 - I0 = zeros(1, Tsteps);
20 - %I0 = I0 + 10*randn(1, Tsteps);
21 - for i=1:15
22 -     I0(1000*i)=9.785*i;
23 - end
24 - %the smallest I0 that created an action potential (so threshold I0) was
25 - %9.785*15= 146.775
26 - % initial conditions:
27 - Volt(1) = -65;
28 - n(1) = 0.318;
29 - m(1) = 0.053;
30 - h(1) = 0.596;

```

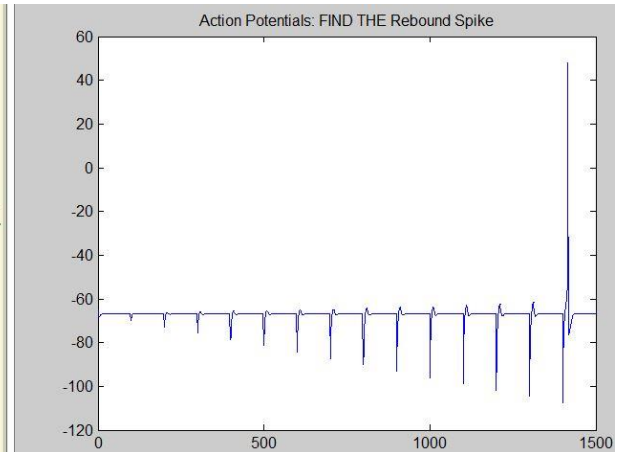


A similar process was used to find the rebound spike inducing stimulus. The I_0 required for a rebound spike was $\sim 15 * (-58) = -870$.

```

6 - T = 1500; dt = 0.05;
7 - Tsteps = round(T/dt);
8 - C = 1; % capacitance (microF/cm^2)
9 - V1 = -77; % leak reversal potential (mV)
10 - g1 = 0.3; % leak conductance (mS/cm^2)
11 - VNa = 55; % Na reversal potential
12 - gNa = 120; % open Na conductance
13 - VK = -77; % K reversal potential
14 - V1 = -61; % leak reversal potential (mV)
15 - gK = 36; % open K conductance
16 - % input current: sets length of applied current to 0 and randomizing
17 - % current amplitude across 0-10 for 4 and 5 comment out the 2nd line and
18 - % redefine current with a vector of currents to find the threshold current.
19 - I0 = zeros(1, Tsteps);
20 - %I0 = I0 + 10*randn(1, Tsteps);
21 - for i=1:15
22 -     I0(2000*i) = -58*i;
23 - end
24 - %the current that produced a rebound spike was ~15*-58= -870
25 - %the smallest I0 that created an action potential (so threshold I0) was
26 - %9.785*15= 146.775
27 - % initial conditions:
28 - Volt(1) = -65;
29 - n(1) = 0.318;
30 - m(1) = 0.053;
31 - h(1) = 0.596;

```



The refractory periods were determined using threshold and an extremely high stimulus current at evenly spaced points in time, and then decreasing the spacing between them. Note that the conversion from time steps (used to vary the period between stimuli) to time in milli seconds is $\text{time} = 0.05 * \text{timesteps}$.

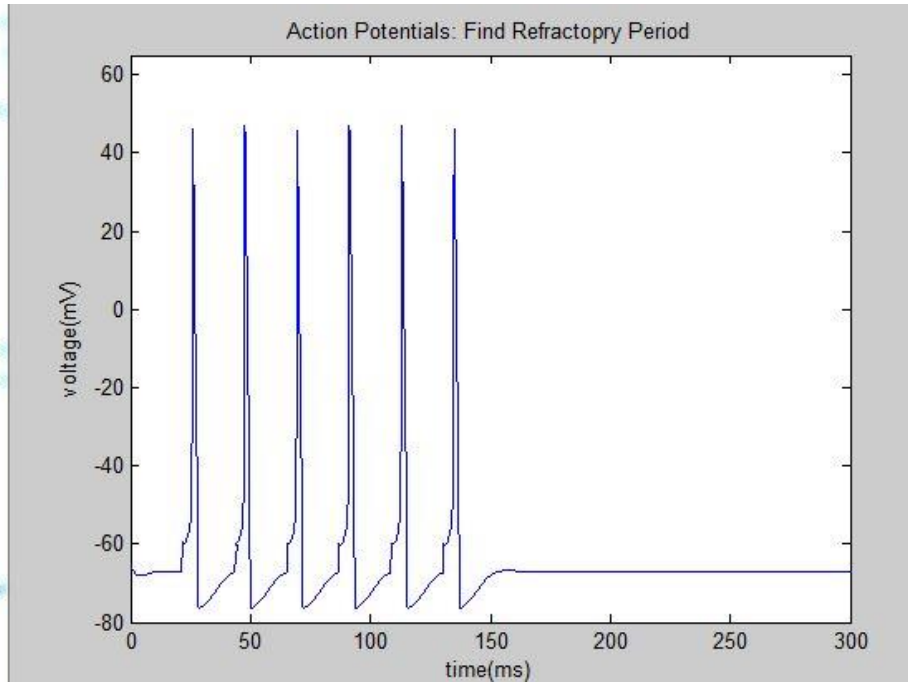
First a value of stimulus current, just above the threshold value that was determined previously, was used ($I = 150$ slightly greater than 146.7705). The stimulus was applied 6 times with decreasing period until every other action potential dropped out.

Here all APs are present with period between stimuli being 435 timesteps and stimuli being 150

```

% current amplitude accord
% redefine current with
I0 = zeros(1, Tsteps);
%I0 = I0 + 10*randn(1, Tsteps);
%for i=1:15
% I0(2000*i) = -58*i;
%end
for i=1:6
    I0(435*i)=150;
end
%the current that produces
%the smallest I0 that causes
%9.785*15= 146.775
% initial conditions:
Volt(1) = -65;
n(1) = 0.318;
m(1) = 0.053;
h(1) = 0.596;
H = [h]; N = [n]; M = [m];
#####
GNav(1)=gNa;

```

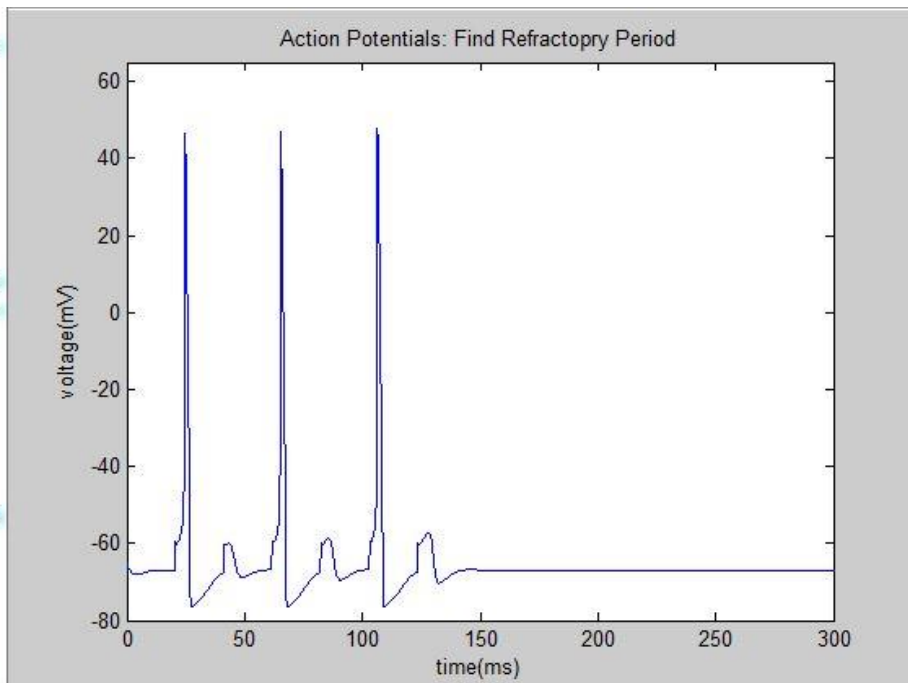


Here they have dropped out when the period between stimuli was dropped to 412 timesteps from 435 timesteps

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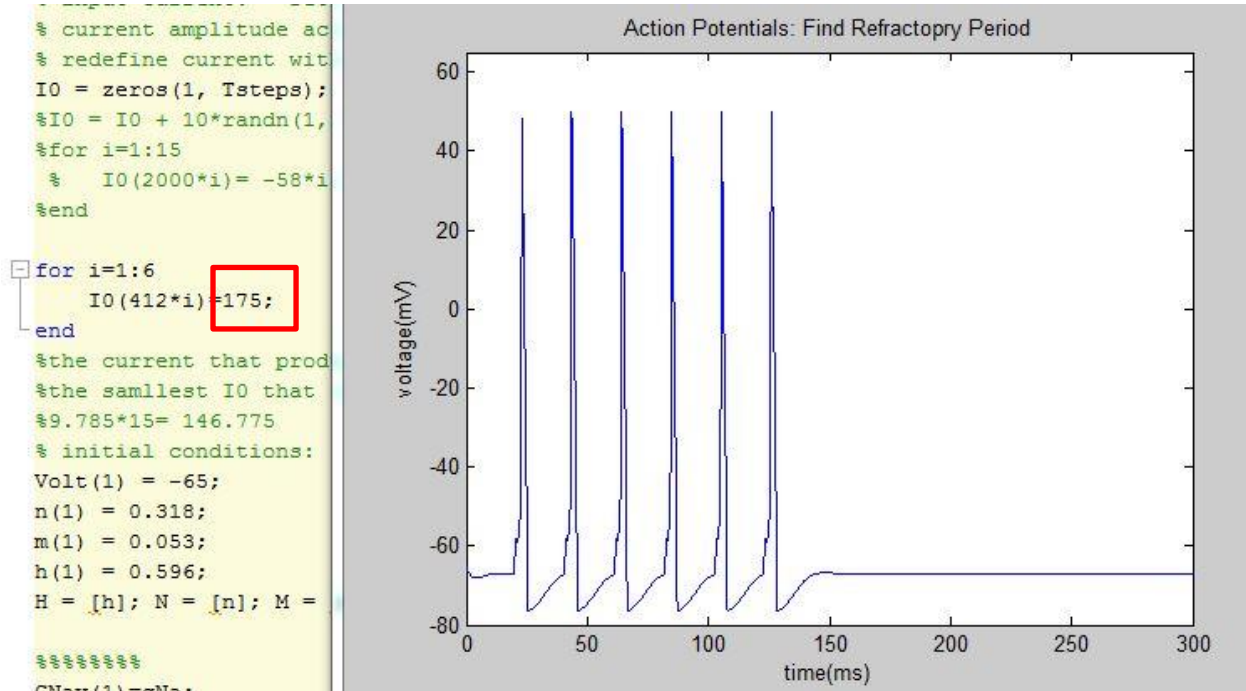
I0 = zeros(1, Tsteps);
%I0 = I0 + 10*randn(1, Tsteps);
%for i=1:15
% I0(2000*i) = -58*i;
%end
for i=1:6
    I0(412*i)=150;
end
%the current that produces
%the smallest I0 that causes
%9.785*15= 146.775
% initial conditions:
Volt(1) = -65;
n(1) = 0.318;
m(1) = 0.053;
h(1) = 0.596;
H = [h]; N = [n]; M = [m];
#####
GNav(1)=gNa;
Gkv(1)=gK;
#####

```

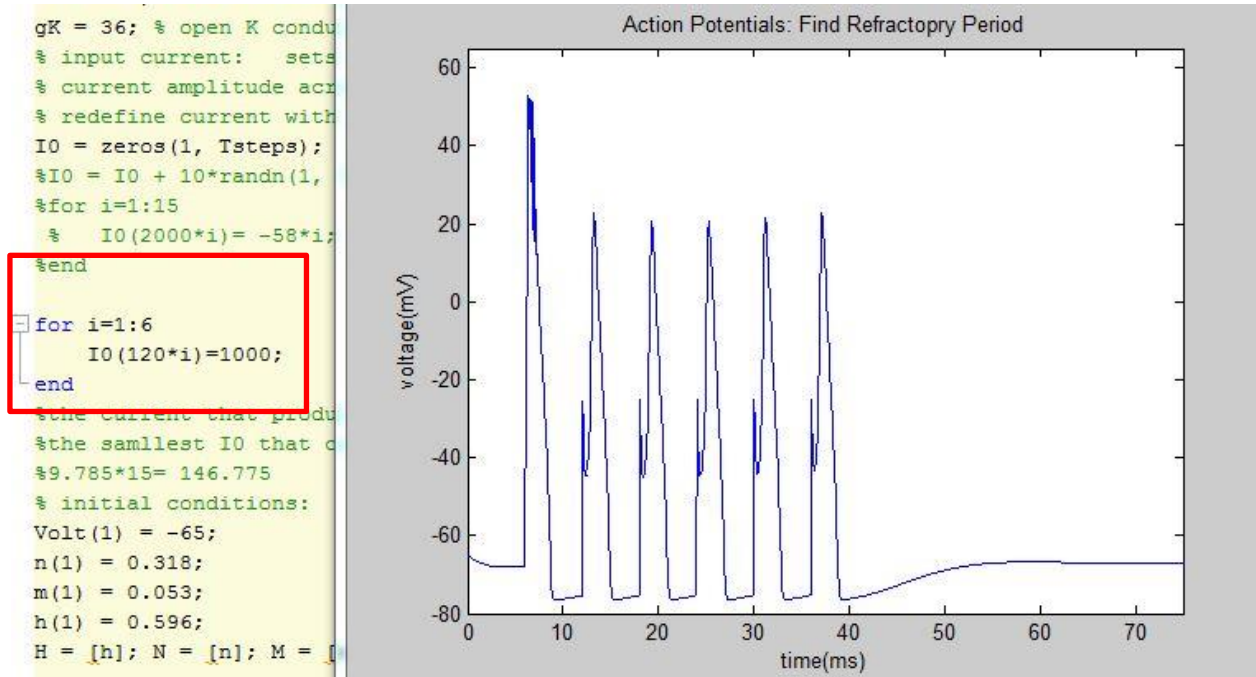


The end of the relative refractory period was ~21ms from the AP that caused it, or between 412&435 time steps. At that refractory period, where every other AP dropped out, increasing the current to 175

brought back the APs that had dropped out, showing that the period was relative refractory, not absolute. This is shown below.



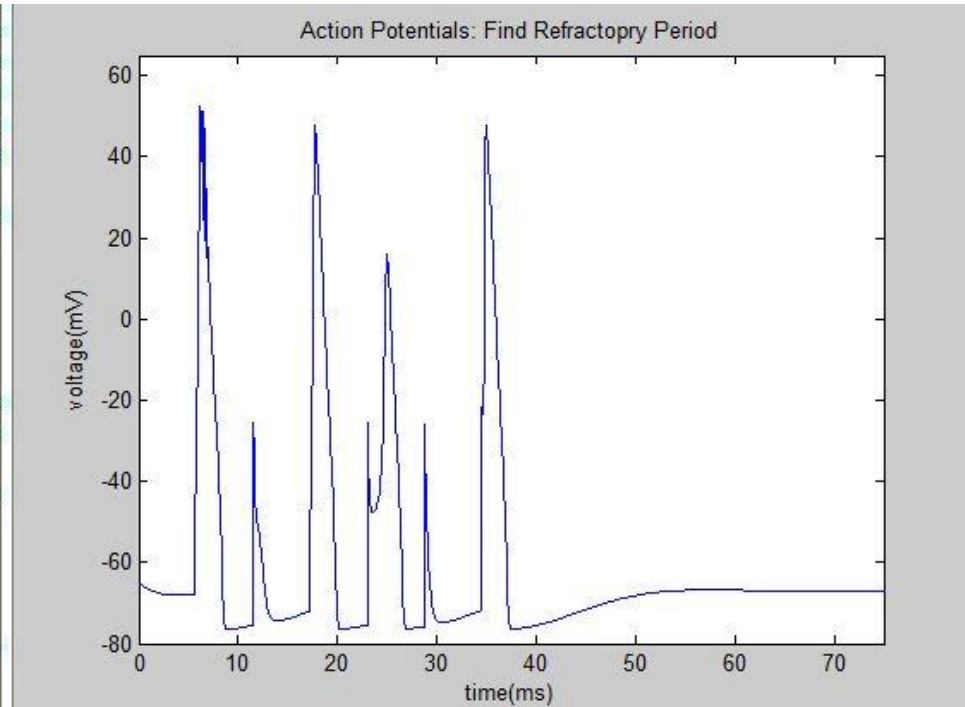
To handle the determination of the absolute refractory period, a much much higher current stimulus was used (1000) to find the absolute refractory period by the same method. This time the absolute refractory was ~5.9ms from the AP that caused it, or between 115&120 time steps.



```

% input current: see
% current amplitude a
% redefine current w
I0 = zeros(1, Tsteps)
%I0 = I0 + 10*randn(2
%for i=1:15
%   I0(2000*i)= -58*
end
for i=1:6
    I0(115*i)=1000;
end
%the current that pro
%the smallest I0 that
%9.785*15= 146.775
% initial conditions:
Volt(1) = -65;
n(1) = 0.318;
m(1) = 0.053;
h(1) = 0.596;
H = [h]; N = [n]; M =
.....

```



The two graphs above show the crossing of the absolute refractory period where even an extremely high stimulus of 1000nA isn't producing full secondary AP because the period between stimuli sinks into the range of the absolute refractory period. The change from being in the relative to being in the absolute refractory periods wasn't an on/off switch. It was fast acting, but showed changes in amplitude as it approached the border.

Discussants: Paras Vora, Jordan Nick, Maeve Woeltje, Matt Everett, Jodi Small, Professor Widder,

References:

1. <http://aries.ucsd.edu/najmabadi/CLASS/MAE140/NOTES/dynamic-2.pdf%E2%80%8E>
2. Koch Chapter 6