

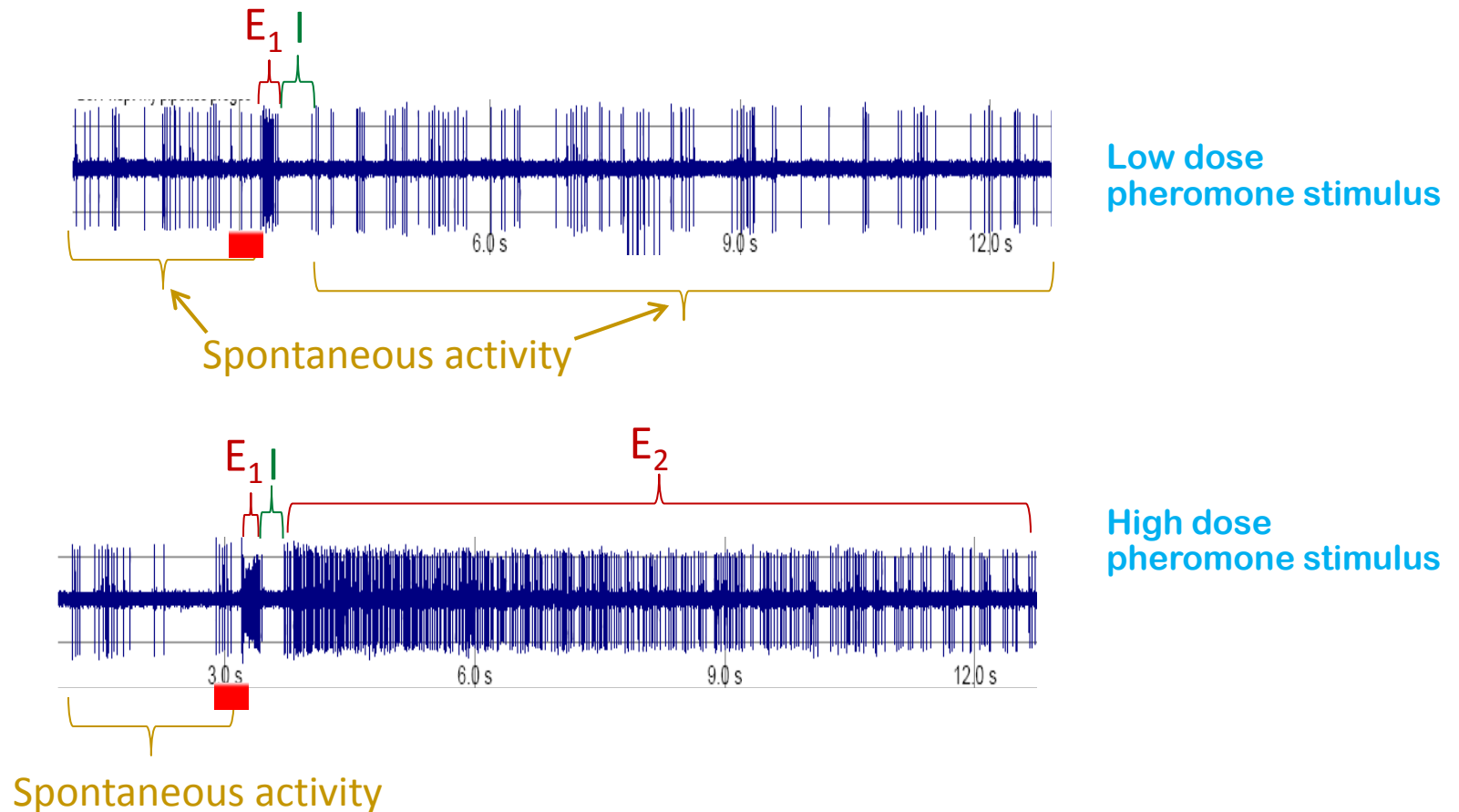
# A BIOPHYSICAL MODEL OF PROJECTION NEURON IN MOTH PHEROMONE CENTRAL PROCESSING SYSTEM—MGC

**Pherosys meeting 2010 INRA Versailles**

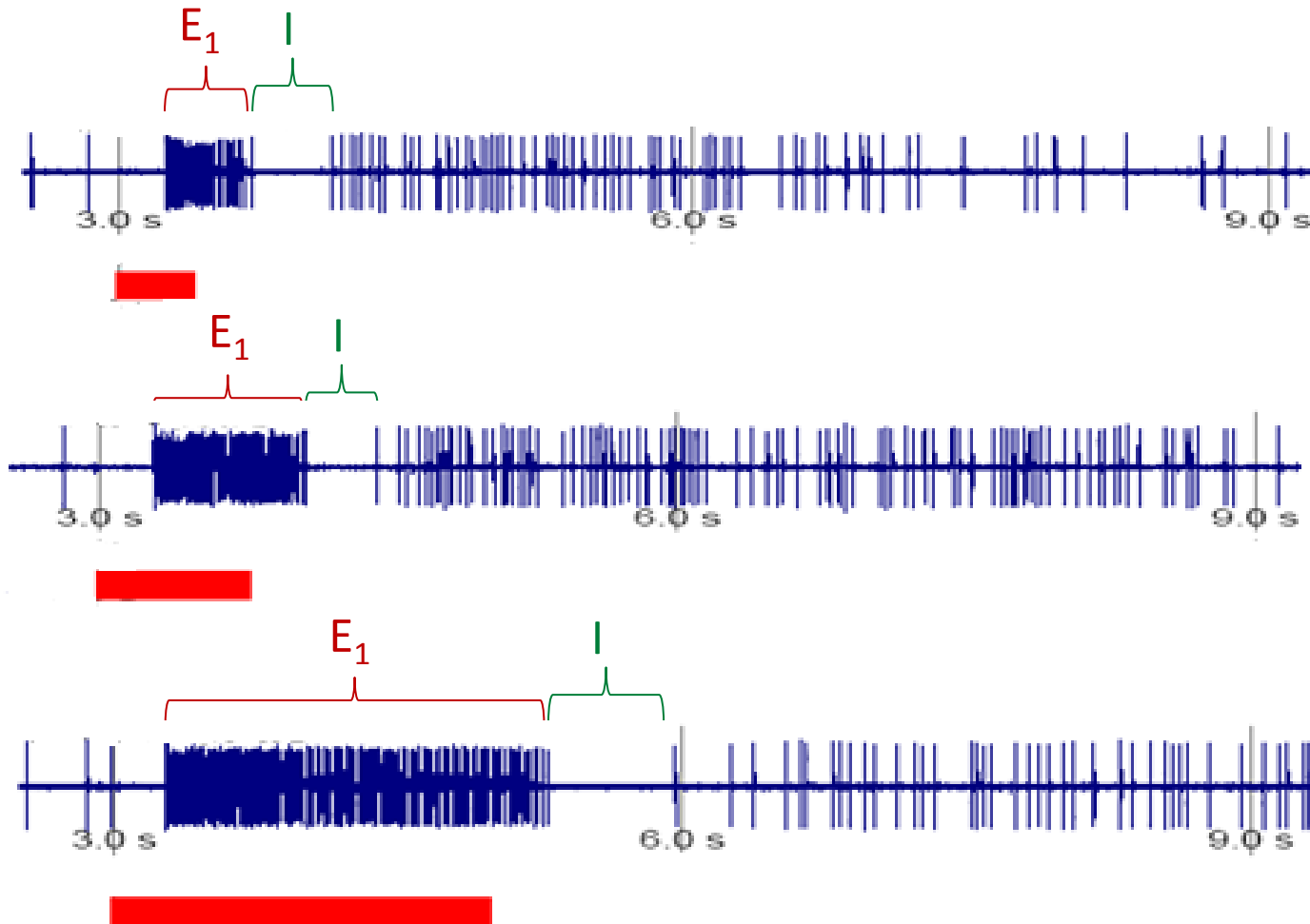
**Yuqiao Gu, Jean-Pierre Rospars, Antoine Chaffiol and Dominique Martinez**

# Experimental data from extra-, intracellular and patch-clamp recordings

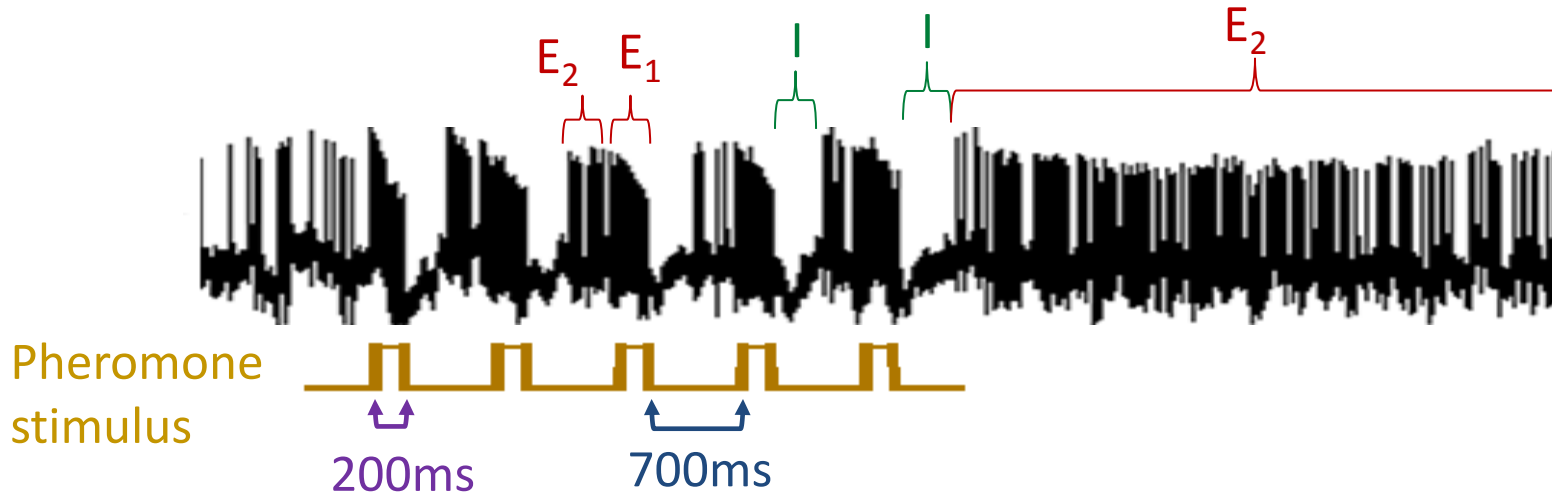
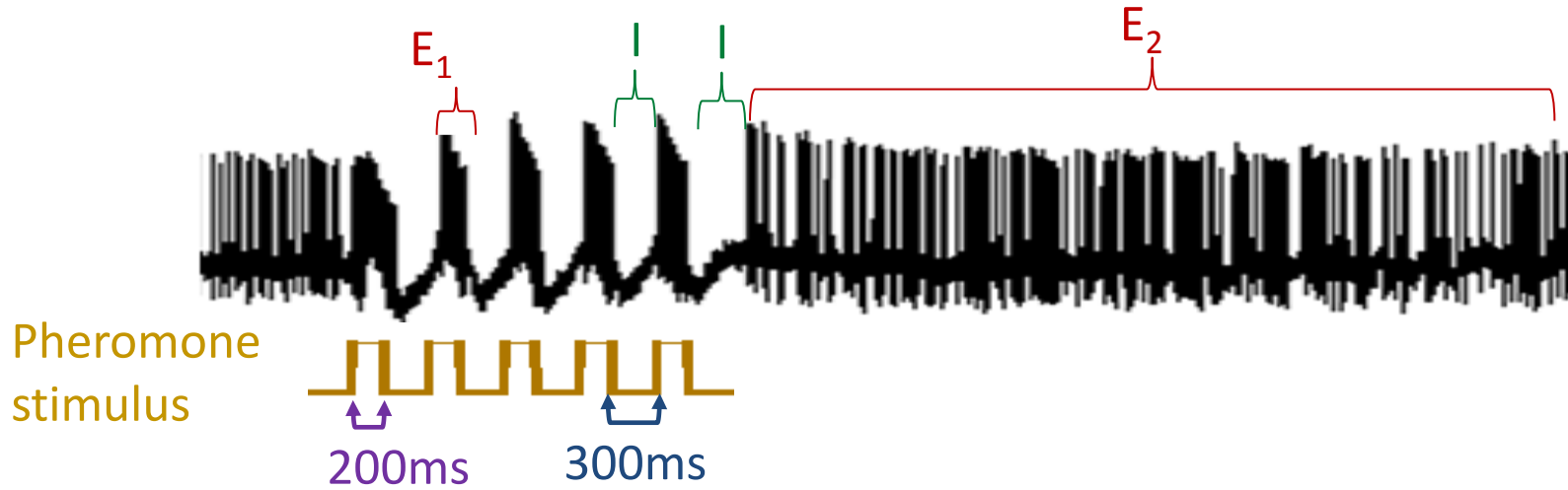
## 1. Extracellular recorded response patterns of the moth pheromone sensitive projection neuron (PN) in MGC



'E<sub>1</sub>' period increases with stimulation period  
while 'I' period keeps almost constant



## 2. Intracellular recorded burst response activity of pheromone sensitive PN to repeated periodic pheromone pulse stimuli



### 3. Whole cell Patch-clamp data on the ionic channels on pheromone sensitive PN and on the other types of neurons in insects

#### *Types of the intrinsic ionic currents on PN*

Using whole cell patch-clamp technique on **PNs in cockroach** (Husch et al., 2009) and **in moth** (Mercer and Hildebrand, 2002) , **five** main components of the inward and outward currents were identified.

#### **Two inward currents:**

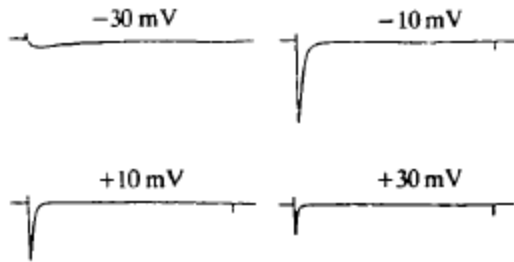
- A fast activating/inactivating  $\text{Na}^+$  current ( $I_{\text{Na}}$ )
- A smaller, slowly inactivating inward  $\text{Ca}^{2+}$  current ( $I_{\text{Ca}}$ ).

#### **Three outward currents:**

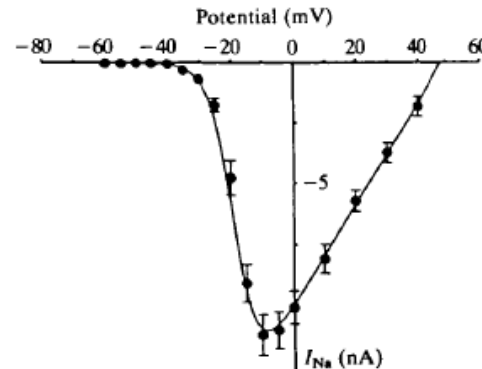
- A transient, voltage-dependent  $\text{K}^+$  current ( $I_{\text{A}}$ );
- A sustained, voltage-dependent  $\text{K}^+$  current ( $I_{\text{K(V)}}$ );
- A  $\text{Ca}^{2+}$ -dependent, outward current  $\text{K}^+$  ( $I_{\text{K(Ca)}}$ ).

## The characteristics of the sodium $\text{Na}^+$ currents $I_{\text{Na}}$

The recorded  $\text{Na}^+$  current  
Traces, that are transient.



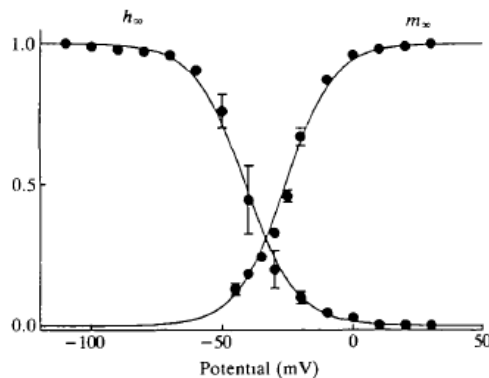
Current-voltage relationship



Lapied et al., 1990

Dorsal unpaired median  
(DUM) cells in cockroach  
*Periplaneta americana*

Voltage-dependence of steady-state  
inactivation and activation

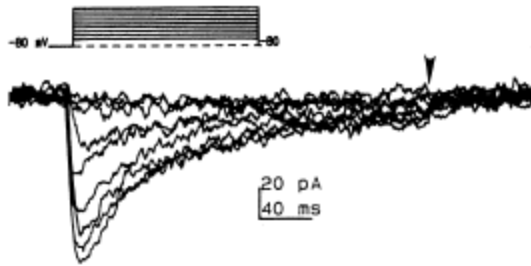


Time constants

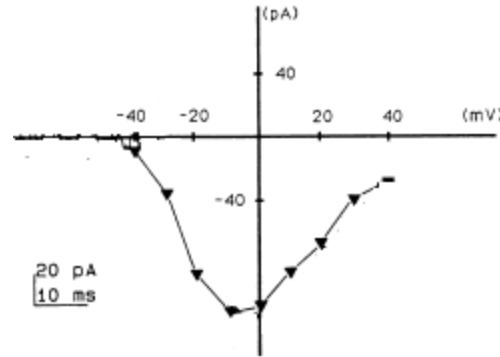
Potential (mV)	$\tau_m$ (ms) <sup>a</sup>	$\tau_h$ (ms) <sup>b</sup>
-25	2.24	6.22±0.89
-20	2.74	2.07±0.54
-15	1.59	1.10±0.09
-10	0.97	1.01±0.02
-5	0.75	0.94±0.04
0	0.59	0.86±0.04
+5	0.52	0.77±0.04
+10	0.45	0.78±0.03

# The characteristics of the calcium $\text{Ca}^{2+}$ currents $I_{\text{Ca}}$

## The recorded $\text{Ca}^{2+}$ currents

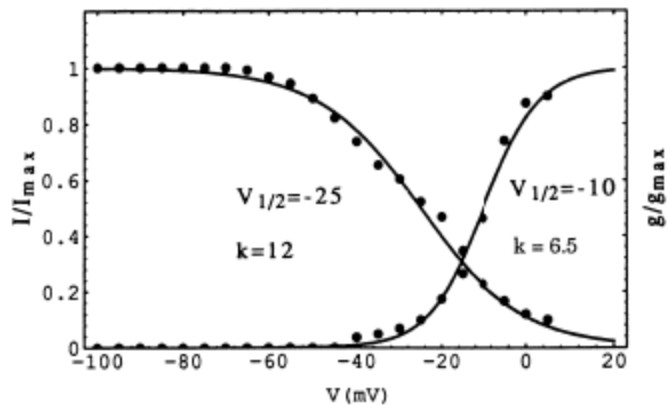


## Current-voltage relationship

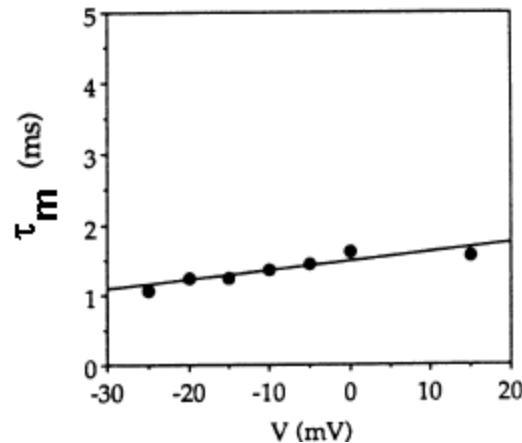


Laurent et al., 1993  
Nonspiking LN in locust

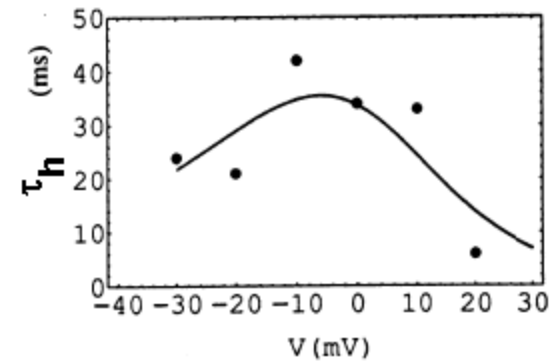
## Voltage-dependence of steady-state inactivation and activation



## Time constant of activation



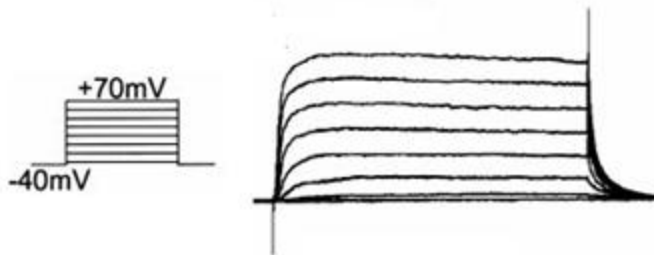
## Time constant of inactivation



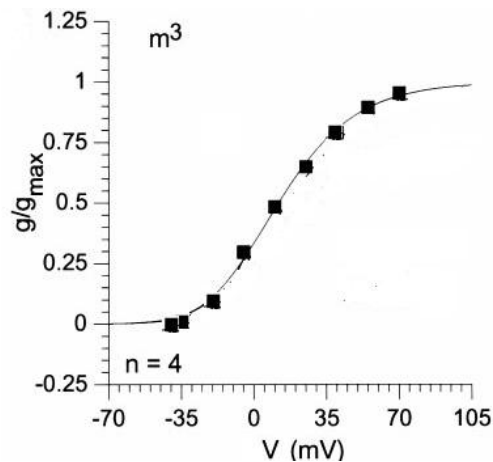
## The characteristics of the voltage-activated sustained potassium $K^+$ currents $I_{K(V)}$

Kloppenborg et al., 1999, in MGC PN of moth *Manduca sexta*

The recorded  $I_{K(V)}$



Voltage-dependence of steady-state activation

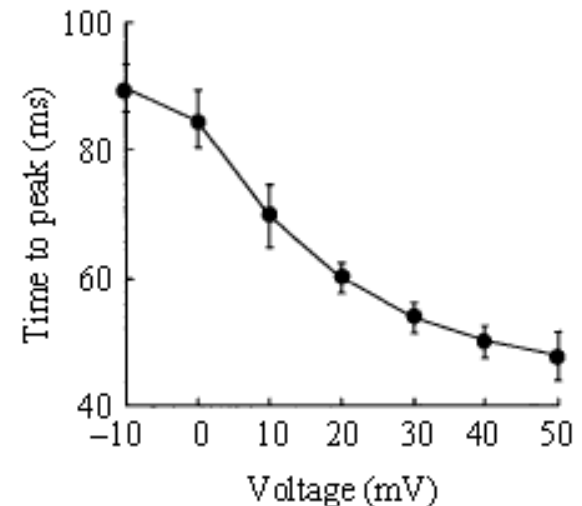


Mercer et al., 1995

In LN in moth

*Manduca sexta*

Time to peak of activation

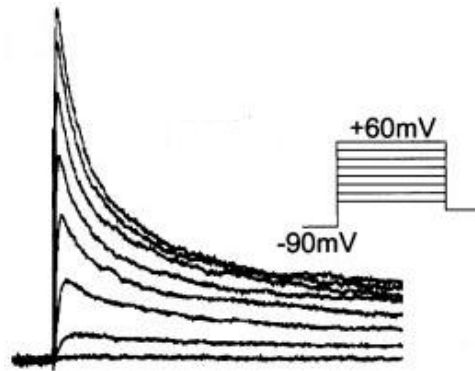




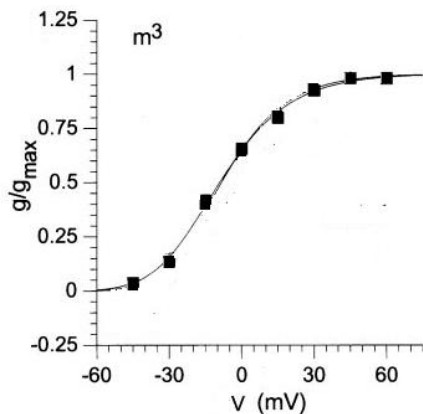
*The characteristics of the transient, voltage-dependent potassium  $K^+$  currents  $I_A$*

Kloppenborg et al., 1999, in PN of moth *Manduca sexta*

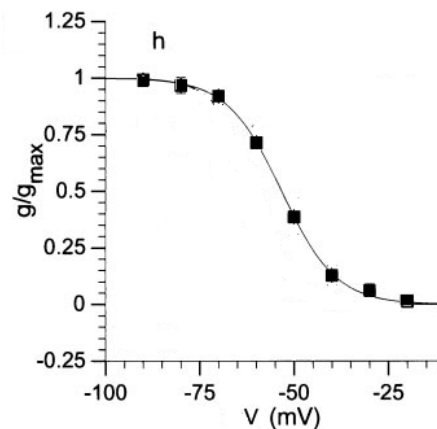
The recorded  $I_A$



*Voltage-dependence of steady-state activation*

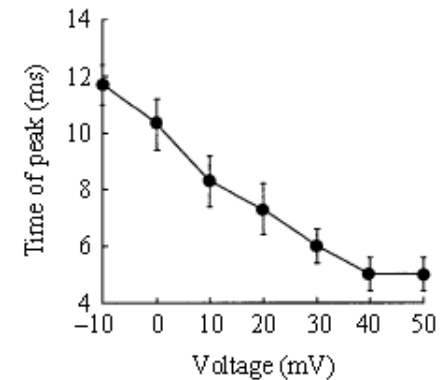


*Voltage-dependence of steady-state inactivation*

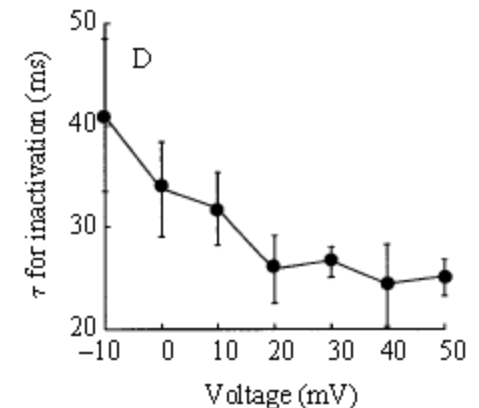


Mercer et al., 1995  
In RR (LN) in moth  
*Manduca sexta*

Time to peak of activation



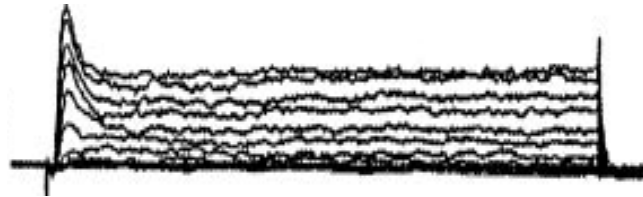
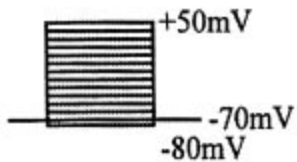
Time constant of inactivation



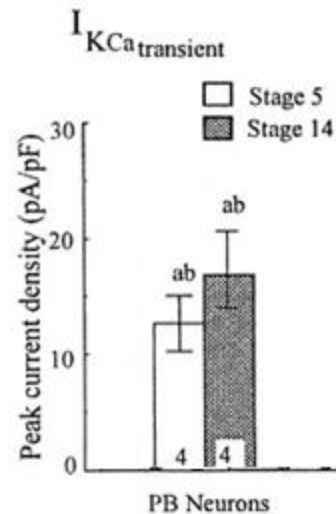
The characteristics of the  $\text{Ca}^{2+}$ -dependent, outward potassium  $K^+$  currents  $I_{K(\text{Ca})}$

Mercer and Hildebrand.,  
2002 In PB (PN) in moth  
*Manduca sexta*

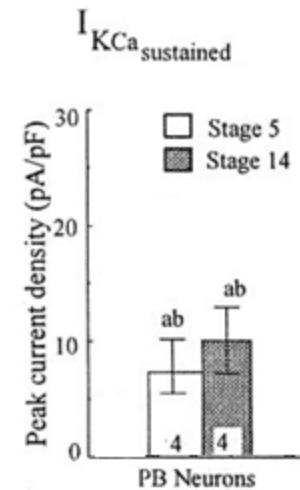
The recorded  $I_{K(\text{Ca})}$  In stage 5 PB (PN)



Comparison of  
mean current  
densities of  
 $I_{K(\text{Ca})\text{transient}}$   
In early (stage 5)  
and late (stage  
14) in PB (PN).



Comparison of  
mean current  
densities of  
 $I_{K(\text{Ca})\text{sustained}}$   
In early (stage 5)  
and late (stage  
14) in PB (PN).



# The biophysical model of the projection neuron (PN)

The differential equation of the membrane potential of the PN

$$C_m \frac{dV}{dt} = -I_{Na} - I_{Ca} - I_{K(V)} - g_L(V - E_L) - I_A - I_{K(Ca)} + I_{inject}$$

$C_m = 22.9$  pF,  $g_L = 11.16$  nS,  $E_L = -61.4$  mV. (PN in cockroach, Husch et al., 2009)

$$I_j = \bar{g}_j m^M h^N (V - E_j)$$

The differential equations of the activation and inactivation variables

$$\dot{m} = (m_\infty - m)/\tau_m \quad \dot{h} = (h_\infty - h)/\tau_h$$

The differential equation of the intracellular  $\text{Ca}^{2+}$  concentration

$$\frac{dCa}{dt} = -f_{Ca} I_{Ca} - (Ca - Ca_\infty)/\tau_{Ca}$$

Mathematical functions describing the voltage dependence of the steady state values and time constants of activation and inactivation variables

$$m_{\infty} = 1 / \{1 + \exp[(V_{0.5act} - V) / s_m]\} \quad h_{\infty} = 1 / \{1 + \exp[(V - V_{0.5inact}) / s_h]\}$$

$$\tau_m(V) = \frac{1}{a_{\tau m, up} e^{(V_{\tau m, 0.5up} - V) / S_{\tau m, up}} + a_{\tau m, dn} e^{(V - V_{\tau m, 0.5dn}) / S_{\tau m, dn}}}$$

$$\tau_{mCa}(V) = 1 + (V + a)b$$

$$\tau_h(V) = \frac{1}{a_{\tau h, up} e^{(V_{\tau h, 0.5up} - V) / S_{\tau h, up}} + a_{\tau h, dn} e^{(V - V_{\tau h, 0.5dn}) / S_{\tau h, dn}}}$$

Mathematical functions describing the  $Ca^{2+}$  dependence of the steady state value and time constant of  $Ca^{2+}$ -dependent  $K^+$  currents

$$m_{\infty} = Ca / (Ca + a)$$

$$\tau_m = b / (Ca + c)$$

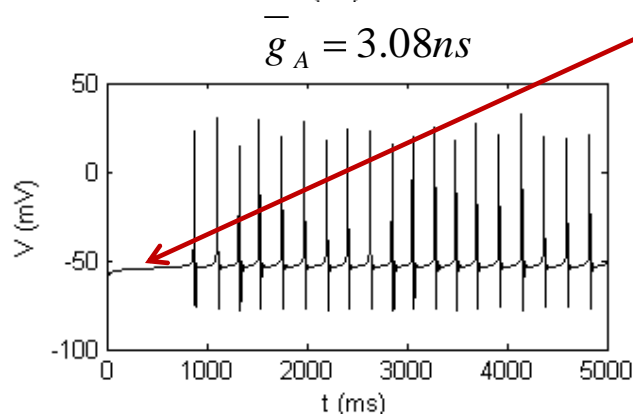
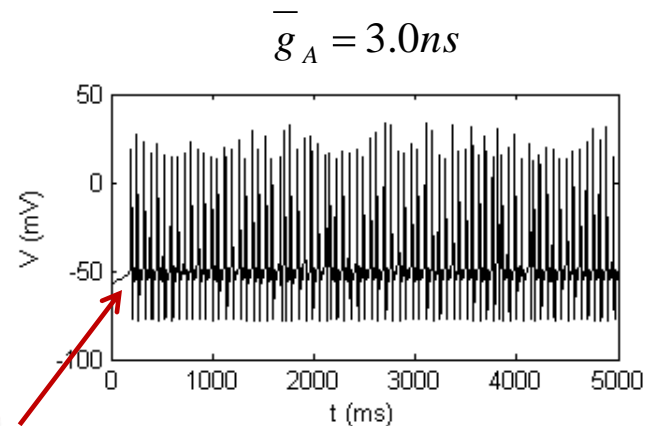
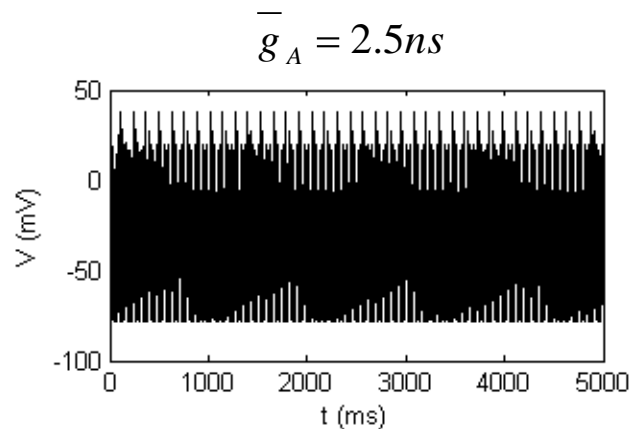
# Preliminary qualitative simulation results

## The intrinsic dynamical properties of the model

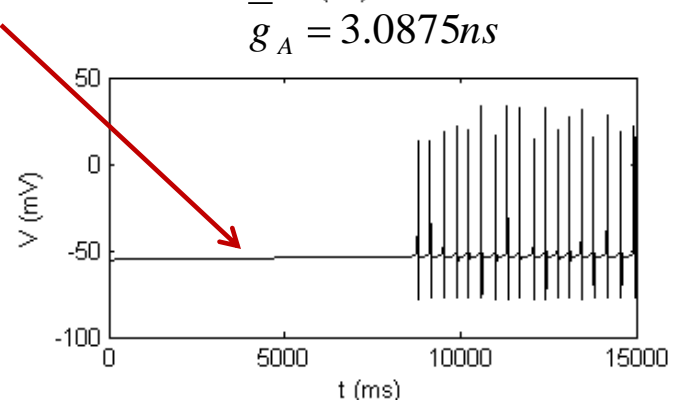
The model PN can be self-excited without any stimulation in certain range of some parameter values. Certain parameters have clear effects on the timing patterns of the action potential

1.  $\bar{g}_A$  of  $I_A$  current strongly affect the starting time and clearly affect firing frequency of the spiking activity

$$\bar{g}_{Kd} = 2.5ns \quad \bar{g}_{KCa} = 0ns$$



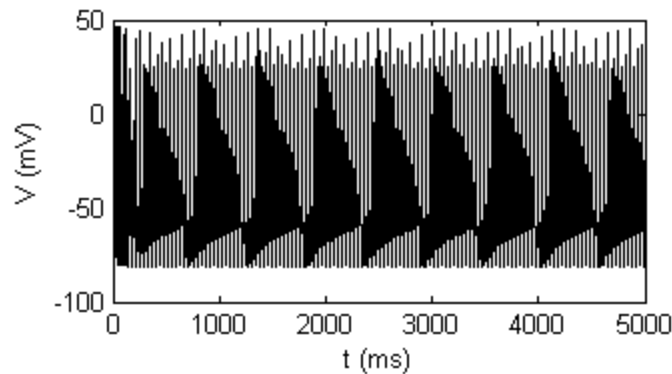
Latency



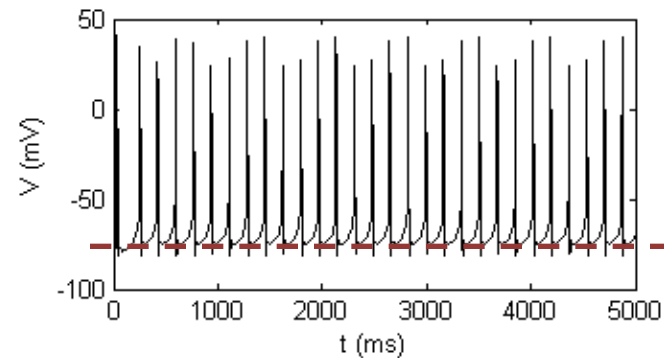
2.  $\bar{g}_{KCa}$  of  $I_{KCa}$  current strongly affect the depth of hyperpolarization and clearly affect the firing frequency of the firing pattern

$$\bar{g}_{Kd} = 2.8ns \quad \bar{g}_A = 1ns \quad \tau_{Ca} = 100ms$$

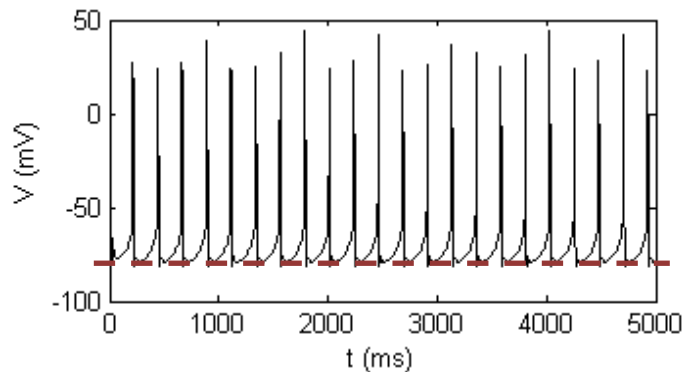
$$\bar{g}_{KCa} = 0.1ns$$



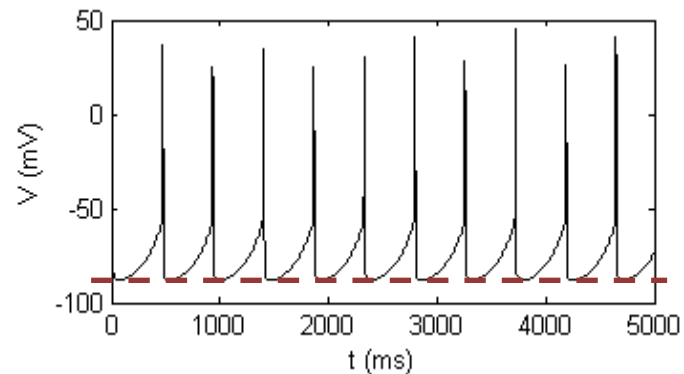
$$\bar{g}_{KCa} = 0.6ns$$



$$\bar{g}_{KCa} = 1ns$$



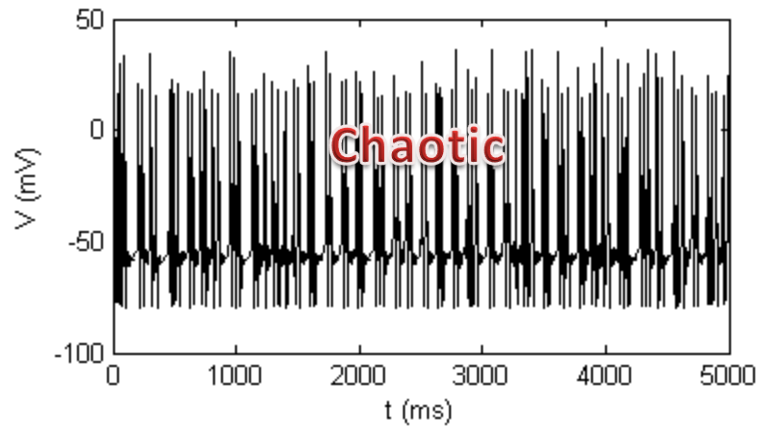
$$\bar{g}_{KCa} = 5.63ns$$



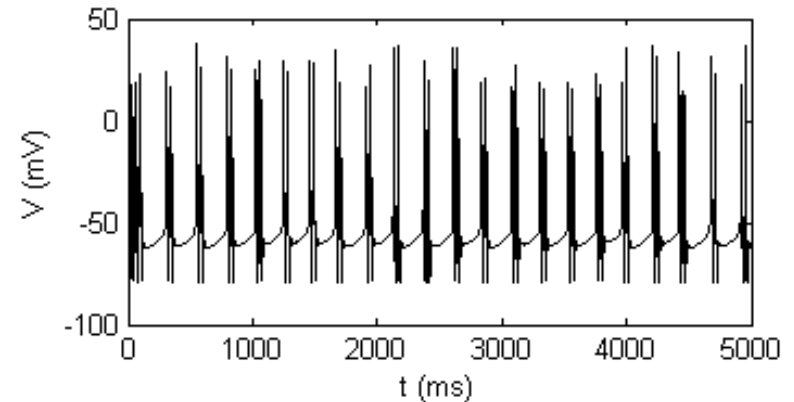
3.  $\tau_{Ca}$  of Ca dynamics strongly affect firing frequency and clearly affect the regularity of the firing pattern

$$\bar{g}_{Kd} = 2.8ns \quad \bar{g}_A = 2ns \quad \bar{g}_{KCa} = 0.1ns$$

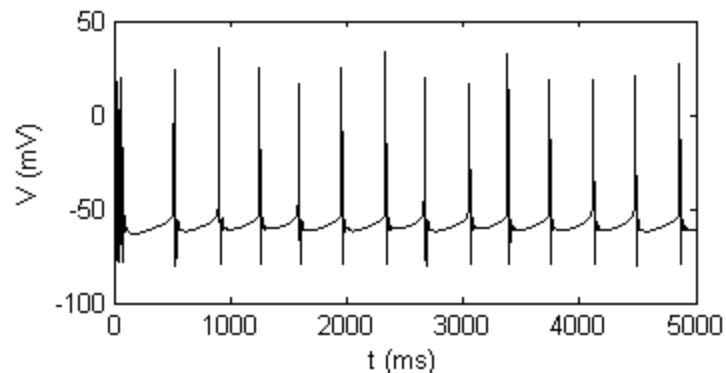
$$\tau_{Ca} = 100ms$$



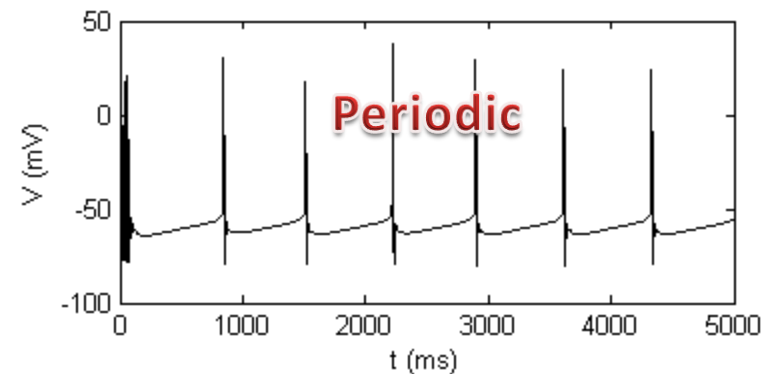
$$\tau_{Ca} = 200ms$$



$$\tau_{Ca} = 500ms$$

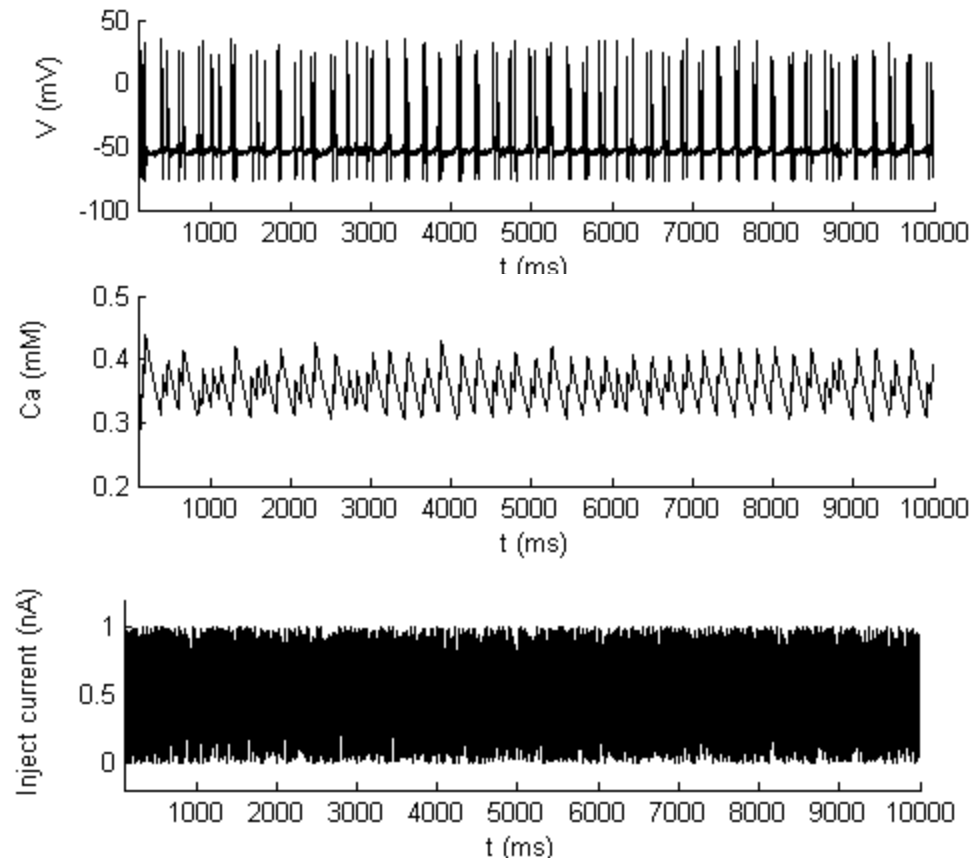


$$\tau_{Ca} = 800ms$$



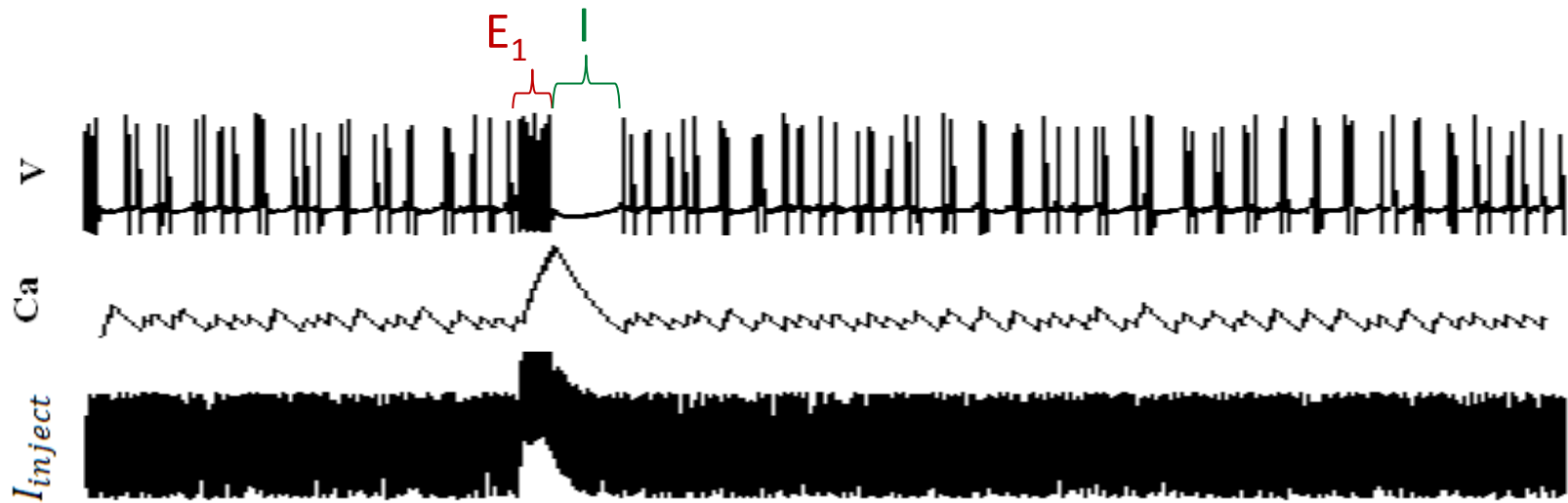
# The generation of spontaneous activity

First, the values of the parameters were adjusted so that model PN cannot fire without any stimulation. Then a noisy current from 0 to 1 nA was applied as  $I_{inject}$



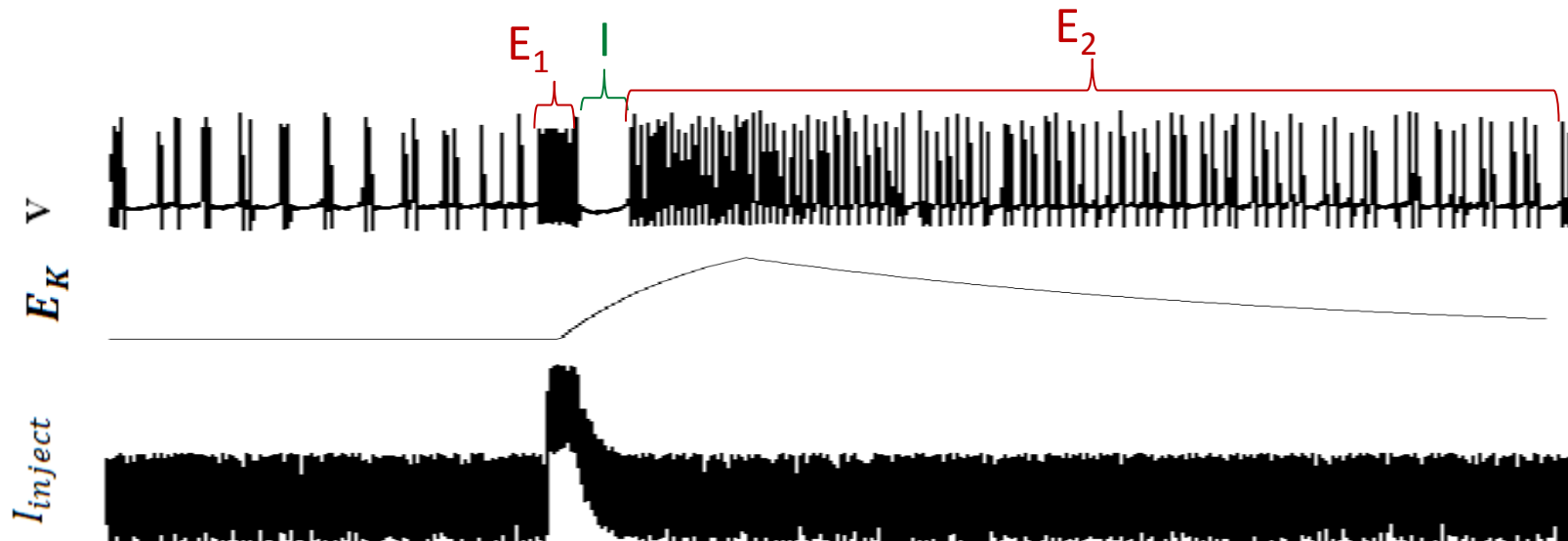


## Reproducing $E_1$ , $I$ and $E_2$ as shown in the experimental data

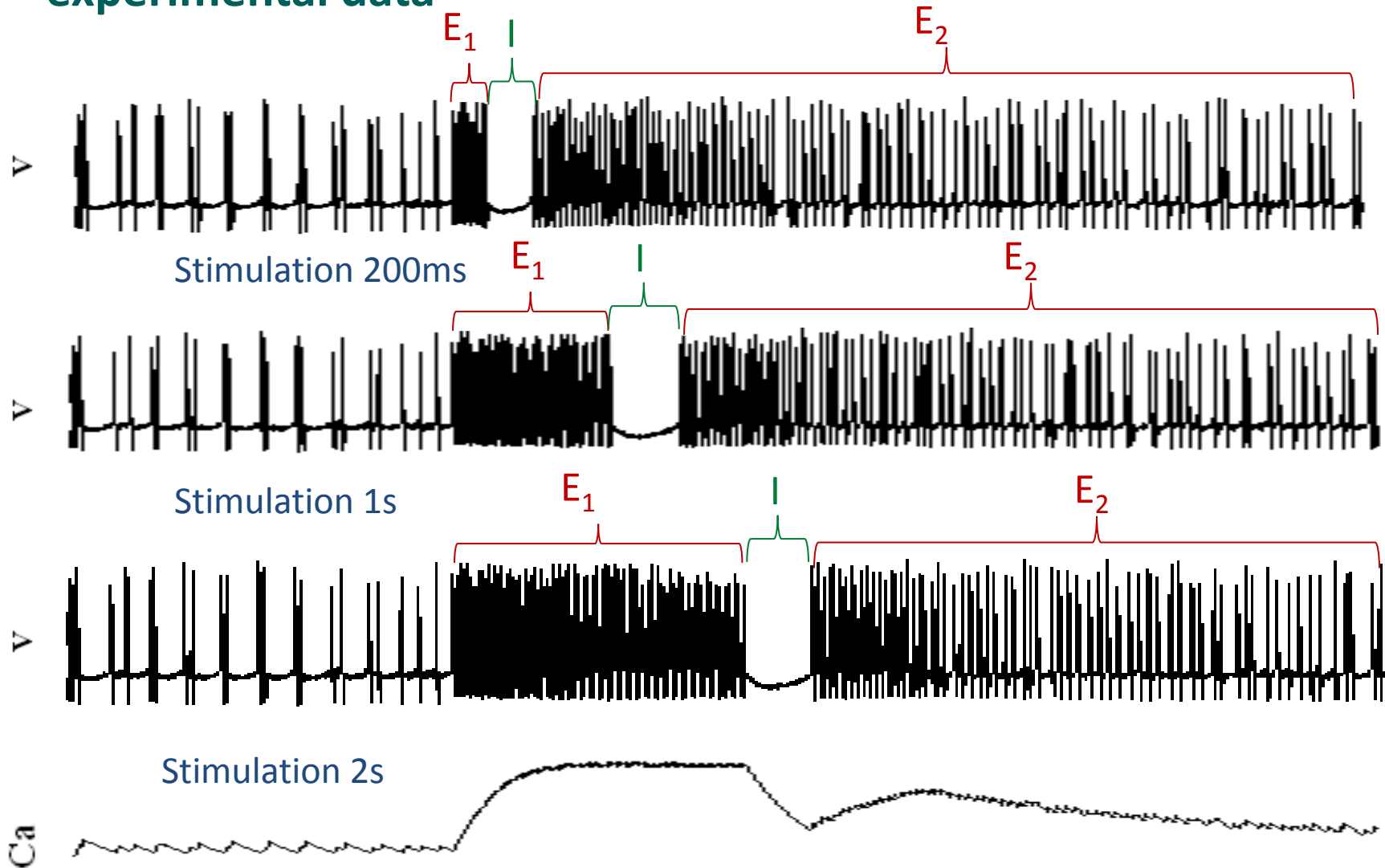


Goriely *et al.*, 2002

Due to the increased dosage of stimuli, extracellular  $K^+$  concentration in the glomerulus increases with the firing of ONR axons. This increases the reversal potential of  $K^+$  currents  $E_K$

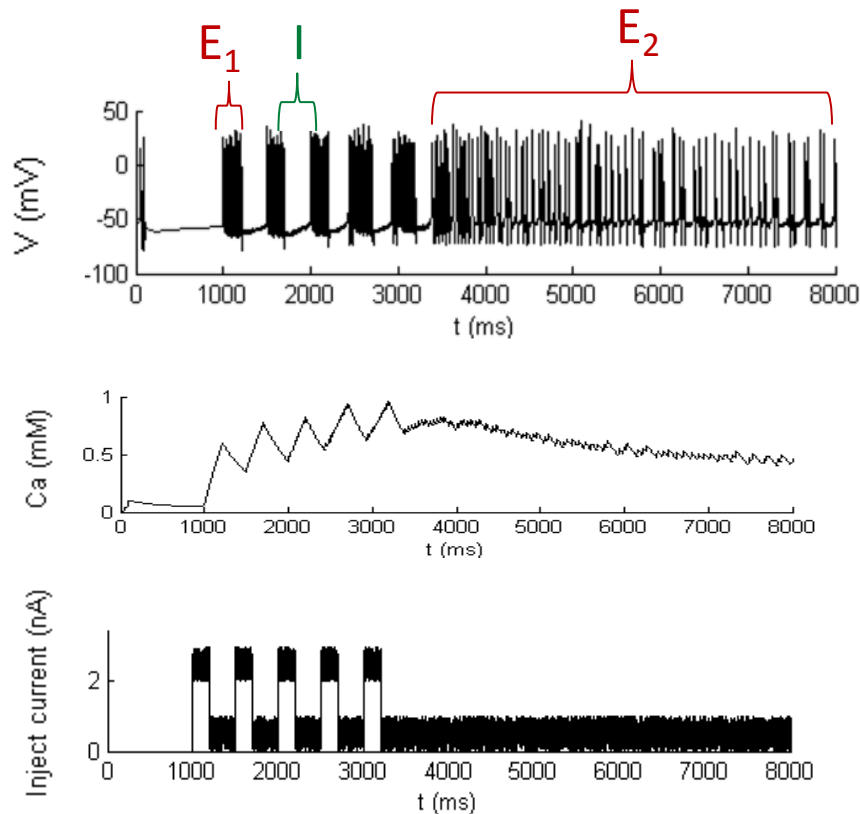


Reproducing that ' $E_1$ ' period increases with stimulation period while ' $I$ ' period keeps almost constant as shown in the experimental data

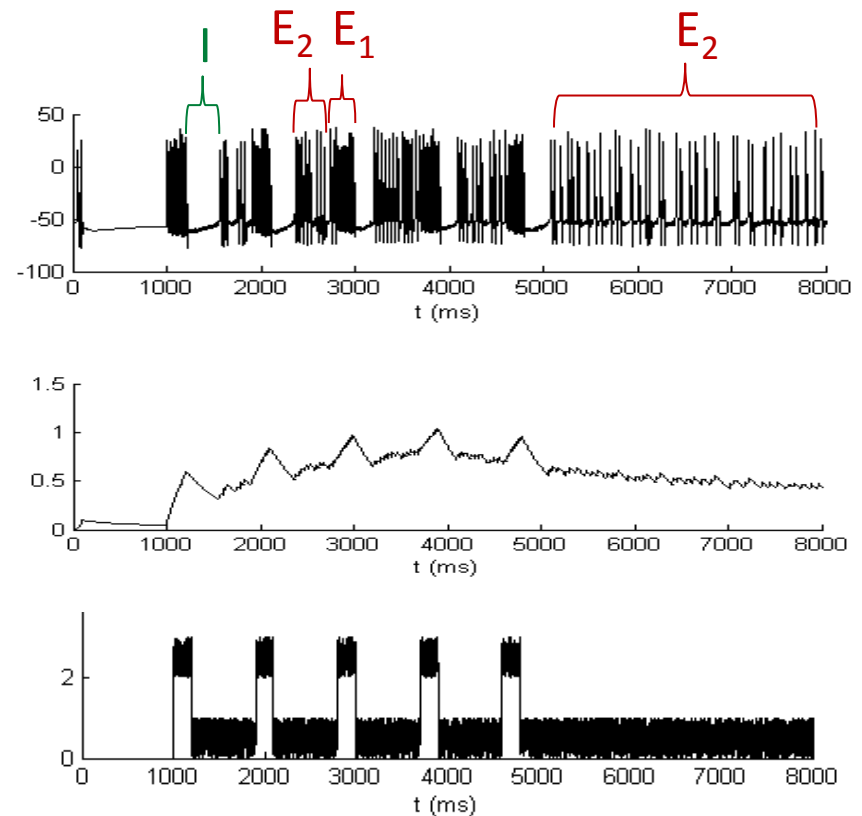


# Reproducing the burst patterns in response to repeated periodic pheromone stimuli as shown in the experimental data

The duration of each pulse is 200ms  
The duration between pulses is 300ms



The duration of each pulse is 200ms  
The duration between pulses is 700ms



# Acknowledgement

**PROF. DOMINIQUE MARTINEZ**

**PROF. JEAN-PIERRE ROSPARS**

**ANTOINE CHAFFIOL**

**Thank you for your attention !**