## A Population Firing Rate Model of Reverberatory Activity in Neuronal Networks

Zofia Koscielniak<sup>1, 2</sup> G. Bard Ermentrout<sup>3</sup>

 Bioengineering and Bioinformatics Summer Institute, Department of Computational Biology, University of Pittsburgh, Pittsburgh, PA 15213
 Carnegie Mellon University, Pittsburgh, PA 15213
 Department of Mathematics, University of Pittsburgh, Pittsburgh, PA

#### **Reverberatory Activity**

- Neural networks are able to maintain persistent activity after temporary inputs
- Input must have sufficient strength
- If so, network will go into longterm stable oscillation



Lau, Pak-Ming, and Guo-Qiang Bi. "Synaptic Mechanisms of Persistent Reverberatory Activity in Neuronal Networks." (2005)

### Fast and Slow Synaptic Activity

#### Fast Activity

- Regulated by neurotransmitter signals
- Result of action potential in adjacent neuron
- Come in short bursts and cause depolarization that is quickly reverted back to resting state
- ~ 5ms

#### Slow Activity

- Regulated by calcium ions
- Elevates potential of membrane slightly
- Takes much longer to return to resting state
- ~200 ms

#### Spike Frequency Adaptation and Synaptic Depression

# Spike Frequency Adaptation Gradual reduction of firing frequency Implicated in habituation

#### Synaptic Depression

- As a cell fires continuously, it does not have enough time after firing to fully recover neurotransmitter level
- Signals it releases become progressively weaker

## Goals

- Create firing rate models of reverberatory activity in neurons incorporating fast synaptic activity, slow synaptic activity, and spike frequency adaptation or synaptic depression
- Analyze models for accuracy and robustness
- Create networks of coupled neurons based on the firing rate models

#### Deriving a Population Firing Rate Model

Total response of post-synaptic cell at time t

$$\alpha(t-t_1) + \ldots + \alpha(t-t_n) = \sum_{j=1}^{n} \alpha(t-t_j)$$

• The total response of all cells becomes

$$I(t) = \int_{0}^{t} \alpha(t-s) (\sum_{j=1}^{n} pr(s=t_{j})) ds$$

Let μ(t) equal the firing rate
 In a population model, the probability equals the firing rate

$$I(t) = \int \alpha(t-s)\mu(s)ds$$

#### Deriving a Population Firing Rate Model

 The firing rate of the post-synaptic cell is determined by the firing patterns of the pre-synaptic cell

$$\mu_{post}(t) = F(I_{post}(t)) = F(\int_0^t \alpha(t-s)\mu_{pre}(s)ds$$

• F is some nonlinear function of inputs • In population models,  $\mu_{pre} = \mu_{post}$ , so

$$\mu(t) = F\left(\int_{0}^{t} \alpha(t-s)\mu(s)ds\right)$$

#### Spike Frequency Adaptation Model

$$F(x) = \sqrt{\frac{x}{1 - e^{-bx}}}$$

tf

 $sa(t) + \mu(t)$ 

ta

SŤ

SS

sa

$$\mu(t) = \sqrt{\frac{(gf * sf + gs * ss - ga * sa - ithr)}{1 - e^{-b(gf * sf + gs * ss - ga * sa - ithr)}}$$

 $ss(t) + \mu(t)(1 - ss(t))$ 

ts

Parameters

- tf=2
- ts=200
- ta=50
- gf=5
- gs=2
- ga=4
- b=8
- ithr=1

#### Spike Frequency Adaptation Model

Model showed stable, long-term oscillation



<sup>-</sup>ast: blue, Slow: yellow, Adaptation: green

Time (ms)

#### Spike Frequency Adaptation Model

 Nullclines show two equilibrium points, one stable and one unstable

Nullclines

Integrated



#### Analysis: Spike Frequency Adaptation Model

Need to make sure oscillation is robust, and not a fluke occurrence
Treat SS as a parameter, conduct bifurcation analysis
SS as parameter should be same as

average of SS in oscillations

#### Analysis: Spike Frequency Adaptation Model



SS (parameter)

Intersection occurs at .438

### Synaptic Depression Model

$$F(x) = \sqrt{\frac{x}{1 - e^{-bx}}}$$

$$\mu(t) = \sqrt{\frac{(gf * sf + gs * ss) * q - ithr}{1 - e^{-b((gf * sf + gs * ss) * q - ithr)}}}$$

$$sf' = \frac{-sf(t) + \mu(t)}{tf}$$

$$ss' = \frac{-ss(t) + as * \mu(t)^{p} * (1 - ss(t))}{ts}$$

$$q' = \frac{1 - q(t) - alpha * q(t) * \mu(t)}{tq}$$

$$rac{F(x)}{F(x)}$$

S

#### Synaptic Depression Model

Model showed stable, long-term oscillation



Time (ms)

#### Synaptic Depression Model

 Nullclines show two equilibrium points, one stable and one unstable

Nullclines

Integrated



#### Analysis: Synaptic Depression Model

Bifurcation diagram with extended Hopf bifurcation

Intersection of average value of SS the parameter and SS the variable



SS (parameter)

Intersection occurs at .890

The Coupling Terms  $sf[1] = \frac{sf[1] + beta * sf[2]}{1 + beta}$  $sf[2...19] = \frac{sf[j] + beta * (sf[j+1] + sf[j-1])}{1 + 2*beta}$  $sf[20] = \frac{sf[20] + beta * sf[19]}{1 + beta}$  $ss[1] = \frac{ss[1] + beta * ss[2]}{1 + beta}$  $ss[2...19] = \frac{ss[j] + beta * (ss[j+1] + ss[j-1])}{1 + 2*beta}$  $ss[20] = \frac{ss[20] + beta * ss[19]}{1 + beta}$ 

Adaptation of Uncoupled Model

$$F(x) = \sqrt{\frac{x}{1 - e^{-bx}}}$$

$$\mu[1..20] = \sqrt{\frac{gf * sf[j] + gs * ss[j] - ga * sa[j] - ithr}{1 - e^{-b(gf * sf[j] + gs * ss[j] - ga * sa[j] - ithr)}}$$

$$sf[1..20]' = \frac{-sf[j] + \mu[j]}{tf}$$

$$ss[1..20]' = \frac{-ss[j] + \mu[j] * (1 - ss[j])}{ts}$$

$$sa[1..20]' = \frac{-sa[j] + \mu[j]}{ta}$$

$$sa[1..20]' = \frac{-sa[j] + \mu[j]}{ta}$$

$$sa[1..20]' = \frac{-sa[j] + \mu[j]}{ta}$$

•Initial conditions: sf[1]=.4, the rest 0



 Synchronization was observed even with very small beta



#### Synaptic Depression Network

Coupling terms same

$$F(x) = \sqrt{\frac{x}{1 - e^{-bx}}}$$

$$\mu[1..20] = \sqrt{\frac{(gf * sf[j] + gs * ss[j]) * q[j] - ithr}{1 - e^{-b((gf * sf[j] + gs * ss[j]) * q[j] - ithr)}}$$

$$sf[1..20]' = \frac{-sf[j] + \mu[j]}{if}$$

$$ss[1..20]' = \frac{-ss[j] + as * \mu[j]^p * (1 - ss[j])}{is}$$

$$q[1..20]' = \frac{1 - q[j] - alpha * q[j] * \mu[j]}{iq}$$

#### Parameters

- beta = .2
- tf=2
- ts=2000
- gf=4
- gs=2.2
- tq=700
- alpha=5
- b=40
- ithr=1.3
- as=40
- ) p=1

#### Synaptic Depression Network

Initial conditions: sf[1]=.5, q[1..20]=1, rest 0



## Conclusions

- Neural activity incorporating fast synaptic activity, slow synaptic activity, spike frequency adaptation and synaptic depression can be modeled with population firing rate models
- These models are accurate and robust
- Network models simulate the activity of multiple systems incorporating these factors

#### References

"Action Potentials." <u>Wikipedia</u>. 19 July 2006 <http://en.wikipedia.org/wiki/Action\_potential>.

Δ

- Benda, Jan, Longtin Andre, and Len Maler. "Spike-Frequency Adaptation Separates Transient Communication Signals From Background Oscillations." <u>The Journal of</u> <u>Neuroscience</u> 25 (2005): 2312-2321.
  - De La Rocha, Jaime, and Nestor Parga. "Short-Term Synaptic Depression Causes a Non-Monotonic Response to Correlated Stimuli." <u>The Journal of Neuroscience</u> 25 (2005): 8416-8431.
- Edelstein-Keshet, Leah. <u>Mathematical Models in Biology</u>. 1st ed. New York: The Random House/Birkhauser Mathematics Series, 1988.
- Ermentrout, G. Bard. "Neural Networks as Spatio-Terminal Pattern-Forming Systems." <u>Reports on Progress in Physics</u> 61 (1998): 353-430.
  - Lau, Pak-Ming, and Guo-Qiang Bi. "Synaptic Mechanisms of Persistent Reverberatory Activity in Neuronal Networks." <u>Proceedings of the National Academy of Sciences of the</u> <u>United States of America</u> 102 (2005): 10333-10338.

## Questions?

