Using Free/Open-source software to analyze breathing in neonatal rat pups

Kathleen Conner, Michael Morikone

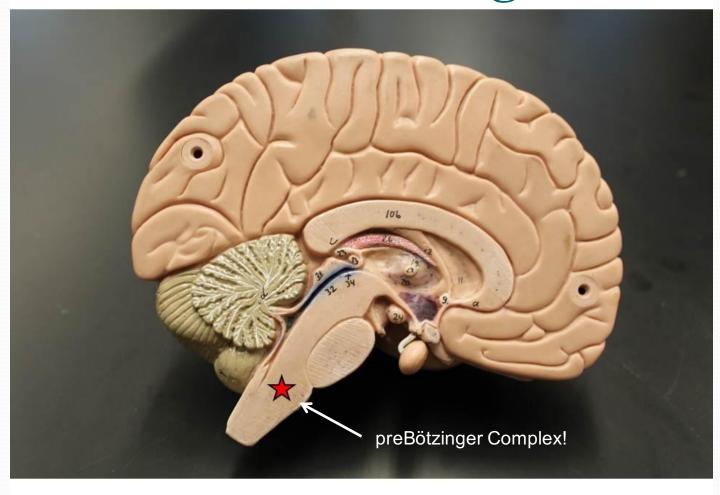


### Background

- In the US and the UK more than 500,000 babies are born prematurely each year
- These babies are more susceptible to infection which often manifests as respiratory problems such as *apnea*, or *periodic breathing*.



# Breathing rhythm originates in the *medulla oblongata*

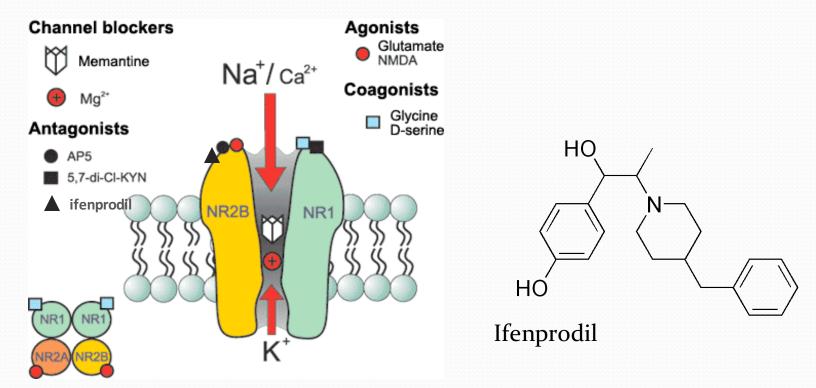


### Background

- Synaptic inputs to nTS neurons—excitatory post-synaptic currents (EPSCs) primarily—are altered by the proinflammatory cytokine, IL-1β.
- Changes in synaptic inputs may be mediated by prostaglandin  $E_2$  receptors but this has not yet been tested.
- To assess changes in synaptic inputs, we developed an analysis tool that allows us to quantify changes in synaptic events (EPSCs).

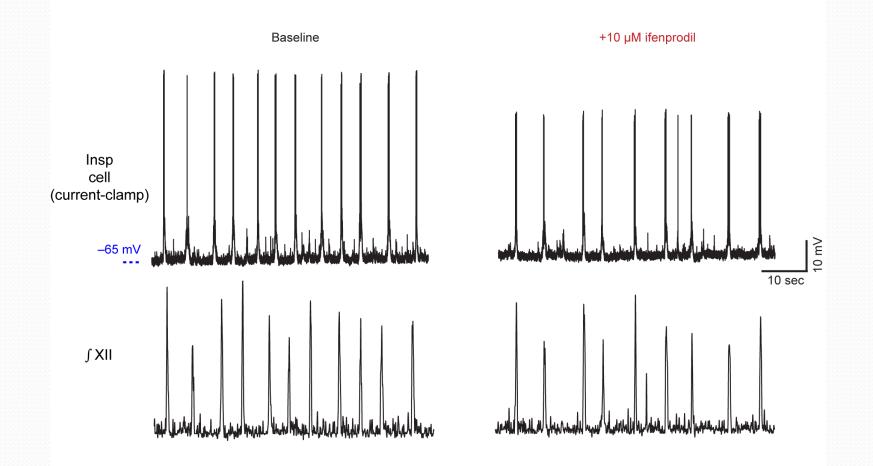
#### NMDA receptors

- NMDA (N-methyl-D-aspartate) receptors are ionotropic glutamate receptors that are non-specific (pass cations)
- NMDA receptors are key excitatory drivers for breathing rhythm



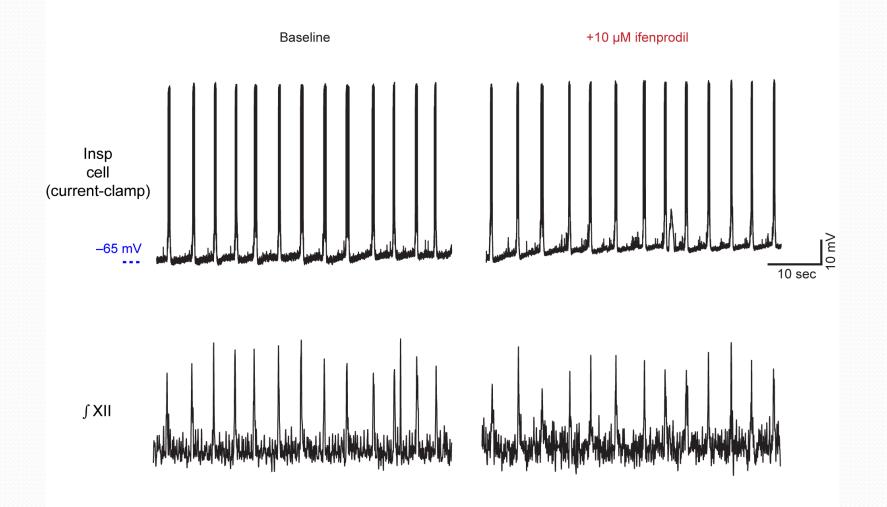


#### Inspiratory neuron from postnatal day 2 (P2) rat





#### Inspiratory neuron from postnatal day 4 (P4) rat



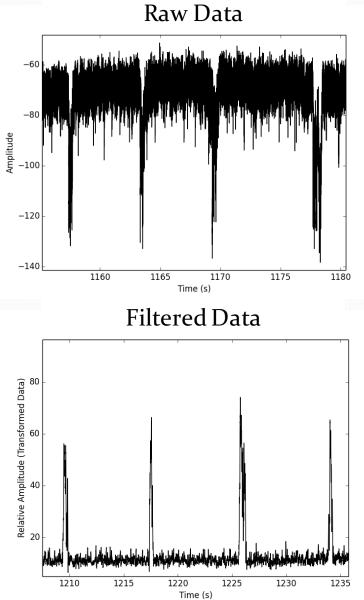
# Hypothesis/Objective

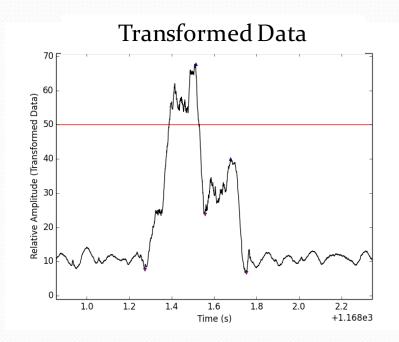
- We hypothesize that neurons in the nTS can alter breathing activity in the presence of the early pro-inflammatory cytokine, IL-1β.
  - This alters the effect of chemosensory inputs to the inspiratory rhythm generator (preBötzinger Complex).
- We hypothesize that NMDA receptors are developmentally regulated, specifically NR2B containing NMDA receptors are present early in development and then are down-regulated.
  - This results in changes in breathing *regularity* and may be a substrate for apneas or periodic breathing in preterm infants!

NTS Hypoglossal (XII) pBC Vagal afferents LPS (IL-1 $\beta$ ) Neurons of the *nucleus* tractus solitarius (nTS) carry Lung Lung afferent traffic from the vagus nerve (X) to the brainstem where autonomic control of the viscera and Diaphragm thermoregulation are located.

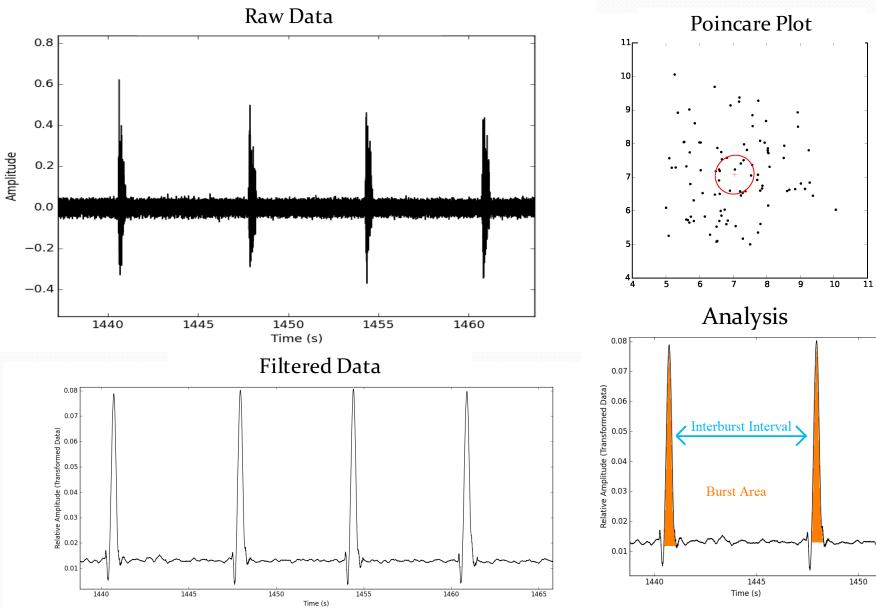
The hypoglossal nerve (XII) innervates the muscles of the tongue which is used in the articulation of speech and swallowing.

#### Raw Data





Raw Data



### What is BASS used for?

- Biomedical Analysis Software Suite (BASS) is a program designed to analyze wave data.
- Event Measurements: Peak amplitude, Peak-Peak Intervals, Duration, Interburst interval, Total Cycle Time, Peaks per Burst, Intraburst Frequency, Burst Area, Attack, Decay.
- Event Detection: Peaks and boundaries (Bursts)

🔄 🔿 🖸 🗋 localhost:8888/notebooks/Desktop/bass-dev/bass-dev/Single%20Wave-%20Interactive.ipynb

 IP[y]: Notebook
 Single Wave- Interactive Last Checkpoint: May 11 14:18 (autosaved)

 File
 Edit
 View
 Insert
 Cell
 Kernel
 Help

 Image: Structure
 Structure
 Cell
 Kernel
 Help

 Image: Structure
 Structure
 Cell
 Cell Toolbar:
 None

#### Welcome to BASS!

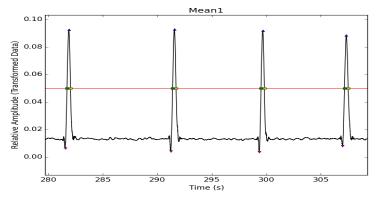
Version: Single Wave- Interactive Notebook.

BASS: Biomedical Analysis Software Suite for event detection and signal processing. Copyright (C) 2015 Abigail Dobyns

This program is free software: you can redistribute it and/or modify it under the terms of the GNU General Public License as published by the Free Software Foundation, either version 3 of the License, or (at your option) any later version.

This program is distributed in the hope that it will be useful, but WITHOUT ANY WARRANTY; without even the implied warranty of MERCHANTABILITY or FITNESS FOR A PARTICULAR PURPOSE. See the GNU General Public License for more details.

You should have received a copy of the GNU General Public License along with this program. If not, see <a href="http://www.gnu.org/licenses/">http://www.gnu.org/licenses/</a>>



0

#### Initalize

Run the following code block to intialize the program. This notebook and bass.py file must be in the same folder.

In [1]: from bass import \*

BASS ready!

#### **Begin User Input**

For help, check out the wiki: Protocol

Or the video tutorial: Coming Soon!

#### Load Data File

Use the following block to change your settings. You must use this block.

Here are some helpful information about the loading settings:

## Advantages of BASS

- Open Source/Free Software
- User friendly
- Consolidation of functions for statistical measurement
- Ease of generation of tables and figures
- Usable with many data sets
- Customizable with programming experience in Python

### So why do all this?

• The purpose of this research is to use free tools to detect changes in breathing regularity and (*hopefully*!) increase the long-term health of preterm babies.



### Acknowledgements

- Christopher G. Wilson, Ph.D., Associate Professor, Center for Perinatal Biology and Dept. of Pediatrics, Loma Linda University
- Abby Dobyns and Lab Technicians at LLUMC
- Arturo Concepcion, Ph.D., Computer Science Department, CSUSB