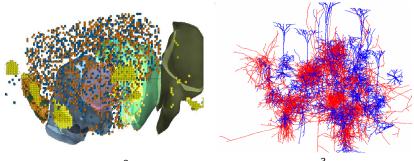
Parallel Collision Detection Over Multi Compartment Morphologies in Python

R. Jarvis Supervisor P. Junor Department of Electronic Engineering La Trobe University May 8, 2016

Introduction

- Made with NEURON-7.4 as a Python module.
- Why did I do this, given the difficulty in staining many adjacent neurons
- Neuromorpho data largely comes from different rodents in different labs.
- Many of the atlas ontologies (neurons) are normalised onto one brain atlas (Allen Brain Atlas)



Motivation

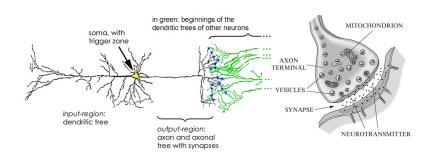
- Light Sheet microscopy, brainbow, and STED may make it possible to digitally recreate more complete forests as opposed to trees.
- Why did I do this? I was inspired by the Blue Brain Project to find out more about real neural networks.
- The Neuromorpho morphologies where/are temporary place holders NeuroMAC is currently able to grow realistic 3D neuron forests in silico, using a forest shapes the trees approach to neural dev.
- Previously Ascoli's L-NAEURON has created a similar tool to grow neurons statistically form is not forest dependant.
- Some models simply distribute Ascoli's in-silico grown trees contiguously in a line on a plane
- Should cortical raster plots have high or low \mathcal{C}_V Chaotic spike trains, may mean that the system has no memory. Regular periodic spike trains may not produce enough unique temporal patterns to be said to be temporal coding.

Motivation

- ullet Spike Distance, graphically showed the C_V as the network activity proceeded through time.
- At the time I thought that what I was doing couldn't be any worse than the small world network topologies assumed by other models.
- I wanted to explore issues such as location dependence of EPSPs, multiplexed rate and temporal codes but instead I looked at simpler questions.
- Networks composed of multicompartment models exhibit much more signal selectivity can involve dendritic subunits can be excited and learn indipendantly.
- Allen Brain has recently acquired gene expression based connectivity maps that may map to selected to the coordinates comprised by SWC files.

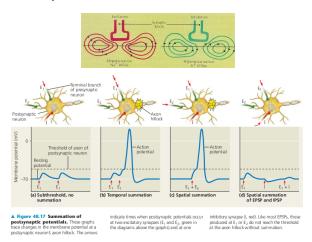
Pre and Post Synapses

 In most what follows I will be referring to pre synaptic locations and post synaptic locations.



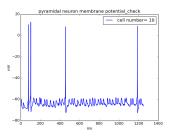
EPSP and **IPSP**

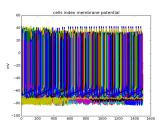
- Some people think of neurons as analog input digital output.
- One or more IPSPs can prevent a spike from occurring, by depressing the membrane potential.

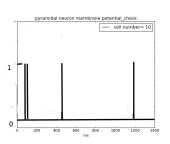


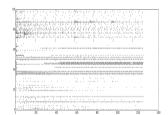
Pre Processing Signals: Spike Trains

• Continuous membrane potential thresholded, the times each neuron fires is stored. Time binning makes signal coarse grained.

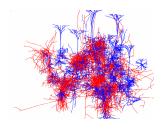








Connectivity and the Adjacency Matrix.



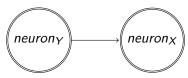
 $I \rightarrow E=2073$, $fb_1 - = 1081$

Information In the Spike Train

- The number of spikes per bin: Δt is the source of uncertainty in the spike train.
- There are two major sources of information:
- 1st the external input (sensory).
- 2nd sources of variability that are intrinsic to the brains dynamics.

An Interpretation of Information Flow:Prediction

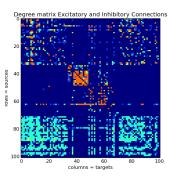
• Using Transfer Entropy we can say that $neuron_Y$ influences $neuron_X$ if $neuron_Y$'s past activity reduces the uncertainty about $neuron_X$'s future activity.

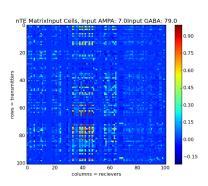


 When a particular type of entropy (uncertaintity) is reduced prediction is increased.

Resulting Information Flow

- One cell can predict another cells activity acting through intermediate neurons
- A reduction in firing probability still represents an information transmission influence.



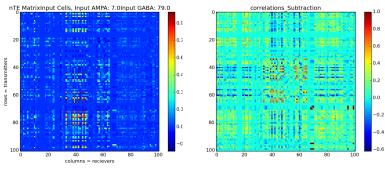


Connectivity

nTE

Information Flow Between Neurons

 Degree matrix shows the quantity and direction of information transmitted between each pair of neurons.

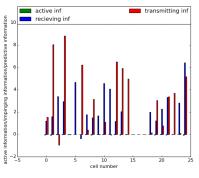


nTE Correlation Matrix

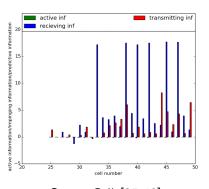
 Presence of information flow contributes validation of reconstruction because when information does flow, it suggests information is not randomly created and destroyed at every node.

Information Flow

 In Von Neuman architecture no conflict between transmitting, translating, and storing information. Respective tasks performed by the bus, the CPU, RAM. One entire neuron simultaneously engages in information transfer, information alteration and information storage.



Sums, Cells[0,24]



Sums, Cells[25,49]

Drawbacks and Future Work

 Predictive changes in spike train information may occur for reasons other than direct connectivity (effective connectivity). Such as when some cells are synchronised to another cell that has a direct connection.

Future work:

- Within cell transfer entropy. The model I implemented contains within cell and between cell feedback. Working memory function of neurons can be inferred from within cell transfer entropy.
- Completely parallelising the network. As well as a partial parallelisation (communication between CPUs), the implementation was performed by formulating the simulation as an embarrassingly parallel problem.
- Inputing a signal into the normalised transfer entropy algorithm that
 has been qauntised into more than two levels, because neural network
 outputs are amplitude modulated signals.

Summary

- Discussed how TE identified functions. TE distinguished information sending neurons from information receiving neurons.
- Transfer Entropy was used to quantify Information Flow within and between neurons.
- I described how information flow in IPSPs can be detected.
- I have discussed how the presence of information flow contributed to the validation of model.
- Generally I have described the analysis of Information flow in a cortical neural network, and why it is of interest.

References



Electrostimulation as a prosthesis for repair of information flow in a computer model of neocortex $IEEE\ 20(2),\ 153-160.$

Neymotin, Samuel A et al (2011)

Synaptic information transfer in computer models of neocortical columns

Journal of computational neuroscience 30(1), 69 - 84.

Gourévitch, Boris and Eggermont, Jos J (2007)

Evaluating information transfer between auditory cortical neurons. *Journal of Neurophysiology* 97(3), 2533 – 2543

Lizier, Joseph T et al (2012)

Local measures of information storage in complex distributed computation

Information Sciences 208, 39-54

The End

Questions?

Trifid

- Trifid is a large Intel system and is VPAC's sixth and largest cluster.
 Trifid has 180 compute nodes, with a total of 2,880 cores. It has a HPL (high performance Linpack) rating is 45.9 TFLOPS. It obviously runs HPC Linux.
- 180 nodes 2,880 cores of Intel E5 − 2670
- 4 GB PC1600 memory per core (64 GB per node),
- In practice could access (99GB on one node)with 6 nodes having 16 GB per core (256 GB per node)
- This computer 4000mbRAM
- 4 nodes will be GPU/MIC capable FDR Infiniband.
- CentOS 6 Linux
- VPAC NFS for home
- VPAC 165TB DDN S2A high performance array for work/scratch/projects (using Lustre).

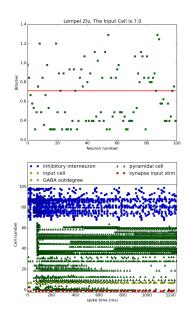
Trifid

 I successfully made an algorithm which could distribute the cells over over the hosts, in NEURON, I did this again in Python.

 Parallelising the wiring diagram was non trivial. It was more complex than necessary using NEURON and MPI alone. Mpi4py has many nice features for easy point to point communication of Numpy Vectors.

Stored Information

stored information



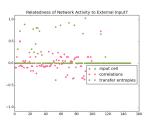
Applications

- Reverse engineering principles of brain function.
- The dynamic clamp Closed loop, real-time NEURON. Models a missing network, and Substites in the missing network behavior with electrode arrays.
- Cogntive neuro prosthesis. If we understand what neurons are doing
 in the hippocampus and prefrontal cortex it is more likely that we will
 be able to replace their functions in the future.
- Neuromorphic Engineering: Noise tolerant AI, by using brain principles, such as computation with noise, using spiking neural networks.
- Epilepsy seizure detection with real time spike distance, in recording electrodes.

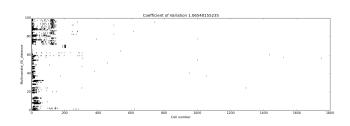
Applications

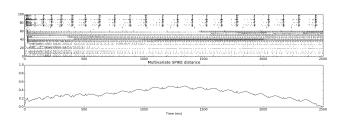
- Reverse engineering principles of brain function.
- The dynamic clamp Closed loop, real-time NEURON. Models a missing network, and Substites in the missing network behavior with electrode arrays.
- Cogntive neuro prosthesis. If we understand what neurons are doing
 in the hippocampus and prefrontal cortex it is more likely that we will
 be able to replace their functions in the future.
- Neuromorphic Engineering: Noise tolerant AI, by using brain principles, such as computation with noise, using spiking neural networks.
- Epilepsy seizure detection with real time spike distance, in recording electrodes.

 The information in the network was also related to the synaptic events that were used to stimulate the network (at two cells only).
 However information in the network was more strongly related to the altered information found across the network.



Information Translation





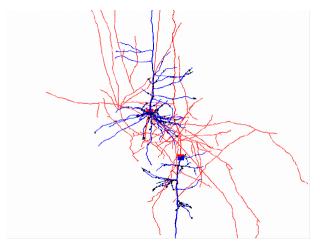
From Reconstructed Cells to Reconstructed Networks.

 In order to move from reconstructed Cells to reconstructed networks, one must connect neurons in a biologically informed way. Spatial and anatomical data from the Allen Rodent Brain Atlas, is used to inform cell centre position and synapse polarity.

 Neurons are connected by conducting an exhaustive search in the neural volume for near contact points and allocating a network connection to the intervening synapse cleft.

From Neurite coordinates to to synaptic cleft

Figure: Two neurons with classified Membrane with Post Synapses



Wiring Algorithm: Searching the volume of spike sources HOC

```
1| proc wire_distances(){
     forsec $01.spk_trig_ls{
     for (x)
         if((x>0)&&(x<1)){
            if (ismembrane("xtra"){
              find_distances(x_xtra(x), y_xtra(x),
    z_xtra(x), secname(), x, $02)
```

```
1//String processing performed elsewhere enabled
    execution of Strings performed here.
    Facilitates network connections.
   sprint(synapse_post, "%s%s%g%s", sec_string2, "
     syn_ = new AMPA(",storex,")")
     execute(synapse_post) //put a post synapse at
     the target
     sprint(synapse_pre, "%s%s%g%s", sec_string1,"
     nc = new NetCon (&v(",num_seg,"),syn_)")
     execute(synapse_pre)
```

Find Contact Points Algorithm

```
forsec $07.spk_rx_ls{
 sec_string1=$s4
 num_seg=$5
 if (strcmp(str_src,str_target) == 0) { break }
    if (Cell[py.int(tail_src)].div.x[py.int(
tail_target)]>3){ break }
      if (ismembrane("xtra")) {
        for(x){
          if((x>0) &&(x<1)){
            r = (sqrt((x_xtra(x) - $1)^2 + (
y_xtra(x) - $2)^2 + (z_xtra(x) - $3)^2)
            if(r<10){//units of um.}
                if (Cell[py.int(tail_src)].div.x
 [py.int(tail_target)] < 3) {
```

Transfer Entropy

The transfer entropy was used to analyse connectivity and to find neurons which were sources of stored information: Then the transfer Entropy (TE) from neuron/process X_1 to neuron/process X_2 is then the amount of reduction of uncertainty between the past of X_1 and the future of X_2 $TE_{X_1 \to X_2} = I\left(X_2^F; X_1^P | X_2^P\right) = H\left(X_1^F | X_2^P\right) - H\left(X_2^F | X_1^P, X_2^P\right)$ By expanding out the equations for entropy H we get:

$$TE_{X_1 \to X_2} = \sum_{k} \sum_{l} \sum_{m} P(X_2^F = k, X_2^P = l, X_1^P = m) \times log_2 \left(\frac{P(X_2^F = k | X_2^P = l, X_1^P = m)}{P(X_2^F = k | X_2^P = l)} \right)$$
(1)

Transfer Entropy

Related to mutual information (the reduction of uncertaintity, when conditional probabilities are considered), except that it involves the difference in entropy between the future state of X and the current state of Y.

By expanding out the equations for entropy H we get:

$$h_{2} - h1 = -\sum_{x_{n+1}, x_{n}, y_{n}} p(x_{n+1}, x_{n}, y_{n}) log_{2} p(x_{n+1} | x_{n})$$

$$+ \sum_{x_{n+1}, x_{n}, y_{n}} p(x_{n+1}, x_{n}, y_{n}) log_{2} p(x_{n+1} | x_{n}, y_{n})$$

$$= -\sum_{x_{n+1}, x_{n}, y_{n}} p(x_{n+1}, x_{n}, y_{n}) log_{2} p(\frac{x_{n+1} | x_{n}, y_{n}}{x_{n+1} | x_{n}})$$

$$h_{2} - h_{1}$$

gives the amount of information that is gained by considering the probability of Y firing and X firing too.

Within Cell Feedback

Within cell feedback

Self projecting synapses

 Back Propagating dendrite action potentials, that can teach post synapses on the dendrite.

Hodgkin Huxley Equations with Synaptic Current

- The Hodgkin Huxley model is a model of spike generation in response to an electrode. The Hodgkin Huxley equation in analytic form is given by the following equation:
- A deterministic equation that first described quiescent conditions.

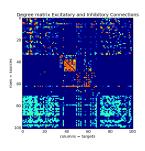
$$\frac{1}{2\pi \mathsf{ar}_i}\frac{\delta^2 V_m}{\delta x^2} =$$

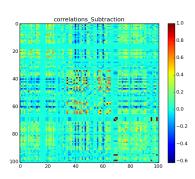
$$C_{m} \frac{\delta V_{m}}{\delta t} + (V_{m} - E_{K}) G_{K}(V_{m}, t) + (V_{m} - E_{Na}) G_{Na}(V_{m}, t) + (V_{m} - E_{L}) G_{L}(V_{m}, t) + I_{syn}$$
 (2)

There are separate equations for the conductance's and that is where the non linearity's emerge from.

ullet Using a similar approach, but with a cubic grid rather than cylinders results in an equation is iterable in discrete temporal steps Δt and spatial steps Δx

$$v_i^{n+1} = v_i^n + \frac{d\Delta t}{4R_i C_m \Delta x^2} (v_{i-1}^n - 2v_i^n + v_{i+1}^n) - \frac{\Delta t G_m}{C_m v_i^n - J_i^n}$$
(3)





Spectral Granger Causality

Spectral Granger Causality was also used.

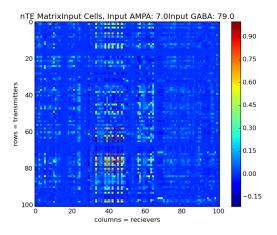


Figure: nTE

Example X_11 : 1110110100110011101101 X_2 : 1010101011011011011

Traditionally in neuroscience the membrane potential has been quantised into two levels, and the time at which a neuron fires is recorded. The motivation codes employed by sensory and motor pathways, are understood to operate on the level of these discrete events.

There is reason to be agnostic about the way information is represented in the brain. One reason is because Action Potentials in many different cortical neurons actually are amplitude modulated.

To counteract this problem there is no reason why the amplitude could not be quantized into ten or more voltage levels. The algorithms which perform entropy calculations are able to accommodate greater than a two symbol alphabet, such that all ten voltage levels could be used to represent different neuron states (different information).

