

IBM Research Division

A Scalable Solution for Specifying and Solving Arbitrary Dense Neural Tissue Model Graphs in a Domain Decomposition Plans and Implications for I/O Bound Applications and Analysis

Presentation to BGAS Workshop, Jülich Supercomputing Centre

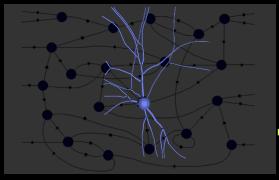
James Kozloski, kozloski@us.ibm.com
T. J. Watson Research Center, Yorktown Heights, NY

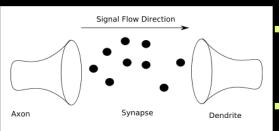


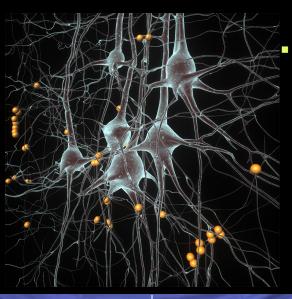
Overview

- Neural Tissue Simulator
- Model Graphs
 - Model Definition
 - Graph Specification
- Scaling
- Neural Current Analyzer



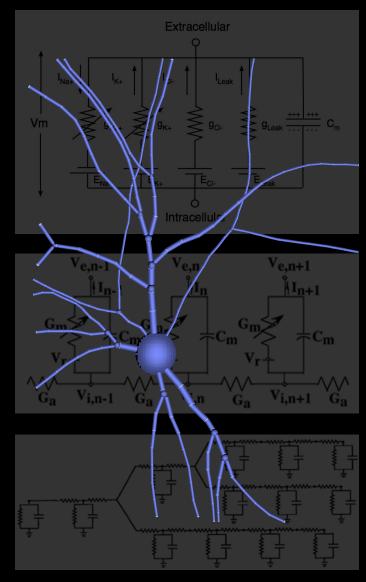






"Neural Tissue Simulation"

- Singe மிர்க் மிர்க்காக models of Hodgkin and Huxley
 - Coupling trans-neuronal
- Muttorcompartment models of single-filters of synaptic release and receptors
 - Multi-compaster gat madels of branched neuronal arbors
- Biological Neural Networks Multi-compartment models of wholeural Parties Sue Simulation





Neural Tissue Simulation

Includes Previous Neural Simulation Constraints

- Replicates a diversity of neuron and synapse types (structural and physiological)
- Uses multi-compartment Hodgkin Huxley models of neurons derived from anatomical reconstructions of real neurons
- Supports synaptic coupling between compartments and attempts to match synaptic distributions from real tissue



Neural Tissue Simulation

Additional Constraints

- 1. Every model in the simulation is embedded within the three-dimensional coordinate system of a neural tissue
- 2. Coordinates for all models are available during initialization and simulation
- 3. Model dependencies, communication, and calculation are some functions of these coordinates

EXAMPLE

 Synapse creation as a function of fiber proximity: Contact Detection



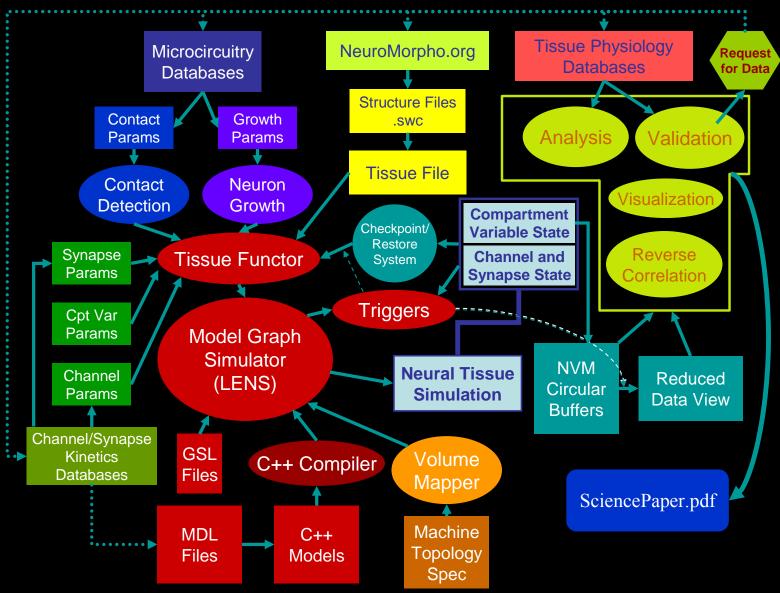
Neural Tissue Simulation

Emerging Opportunities

Model large-scale diffuse tissue phenomena:

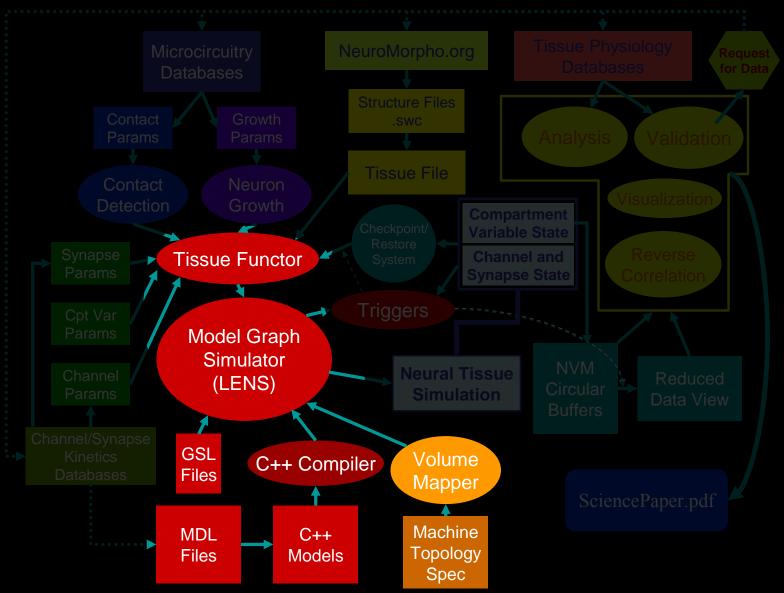
- Widespread gap junctional coupling
- Neuromodulation/plasticity
- Tissue/circuit development
- Tissue/circuit pharmacology
- Brain injury/stimulation
- EEG/BOLD signals

Neural Tissue Simulation Workflow





Neural Tissue Simulation Workflow





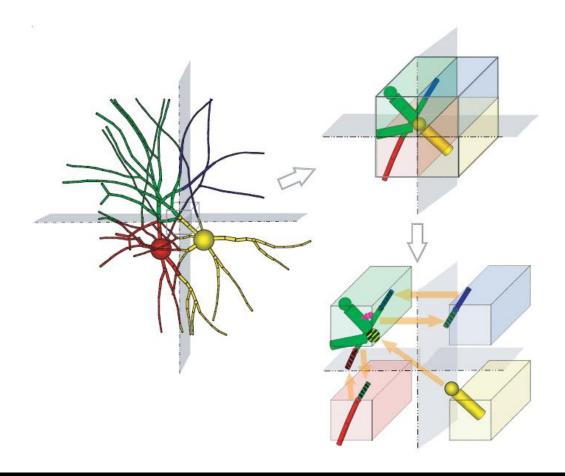
Model Graph Simulator: Infrastructure

Model Definition/ Graph Specification Architectural Overview

- Language for expressing model state, computational phases, communicated state, and model interfaces (MDL)
- Language for composing arbitrary parameterized graphs (GSL)
- Automatic partitioning into work units for multi-threaded execution (SMP)
- •Dynamically constructed, simulation-specific MPI collective communication for multi-process execution of computational phases (MPP)



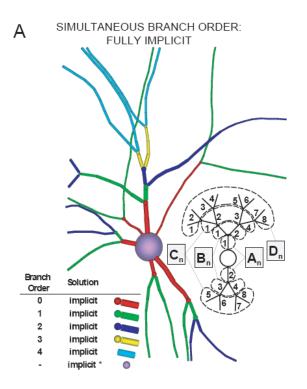
Model Graph Simulator: Tissue Volume Decomposition

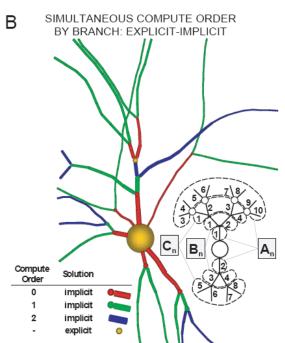


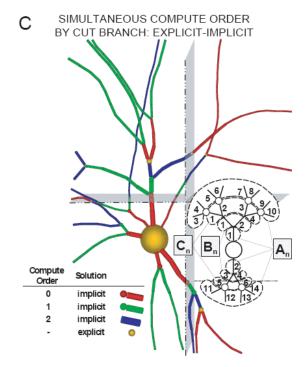


Novel Numerical Approach

- John Wagner, Manager IBM Research Australia / Computational Biology Co-laboratory









Model Definition: Phases



Time Step

Multiphase Algorithm





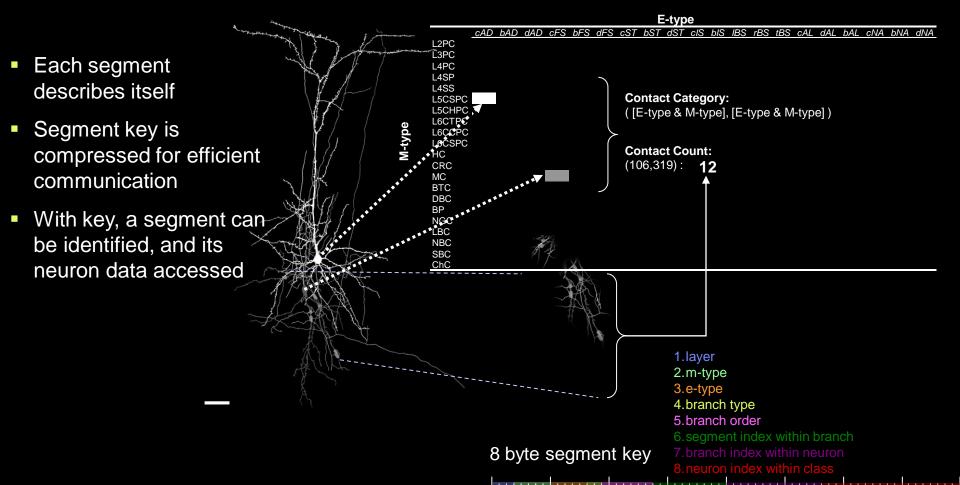
Model Definition: Interfaces



```
Node NaChannel Implements ConductanceArrayProducer, ReversalPotentialProducer
 double [] m;
 double [] h;
 double [] g;
 double [] gbar;
 double []* V;
 Connection Pre Node (PSet.identifier=="compartment") Expects VoltageArrayProducer
  VoltageArrayProducer.voltageArray >> V;
 Connection Pre Node (PSet.identifier=="IC") Expects NaConcentrationProducer {
  NaConcentrationProducer.Na >> Shared.Na_IC;
```

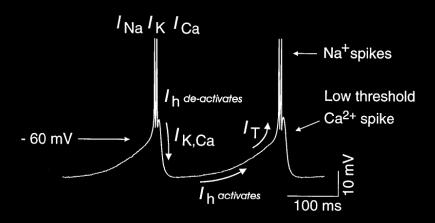


Graph Specification: Key Component Identities

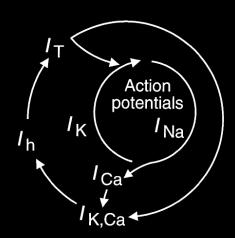




Modeling Objective: Inferior Olive



- Oscillations generated by intrinsic interplay between membrane currents
- Subthreshold oscillations are not driven by spike *input*, but instead constrain and drive spike *output*

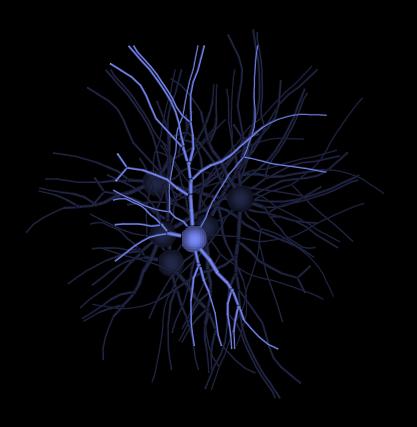


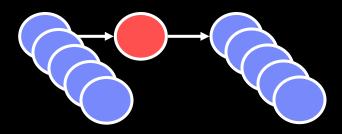


T. Bal and D. McCormick, "Synchronized oscillations in the Inferior Olive are controlled by the hyperpolarization-activated cation current I_h ", J. Neurophysiol. 77:3145-3156, 1997.



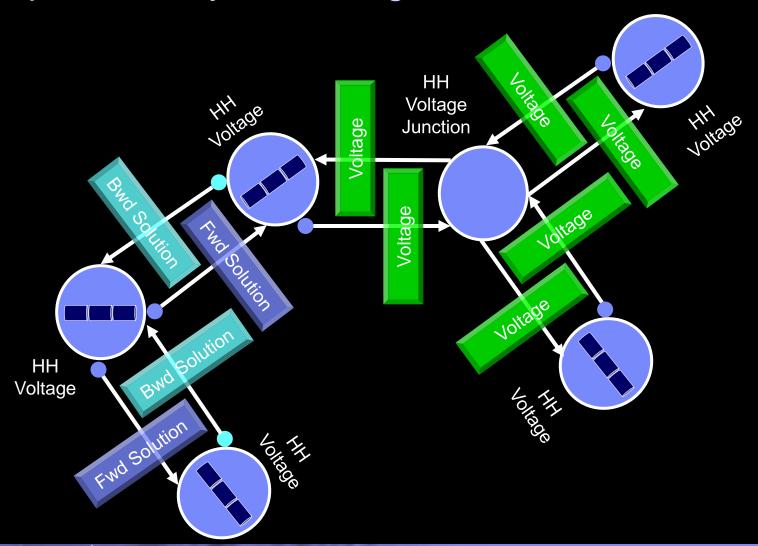
Modeling Calcium Dynamics in IO neurons





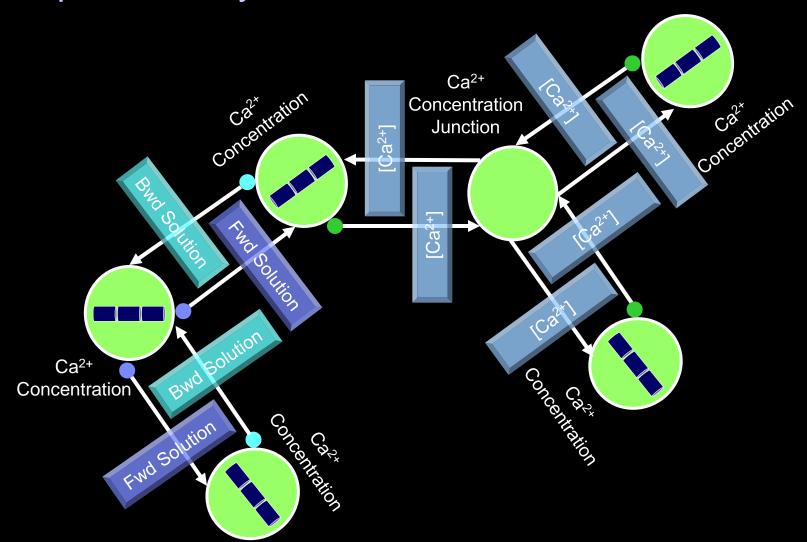


Graph View: Hybrid Voltage Solver





Graph View: Hybrid Calcium Solver





Graph Specification: Compartment Variables

COMPARTMENT_VARIABLE_TARGETS 4 BRANCHTYPE

- O Voltage, Calcium
- 1 Voltage
- 2 Voltage, Calcium
- 3 Voltage, Calcium

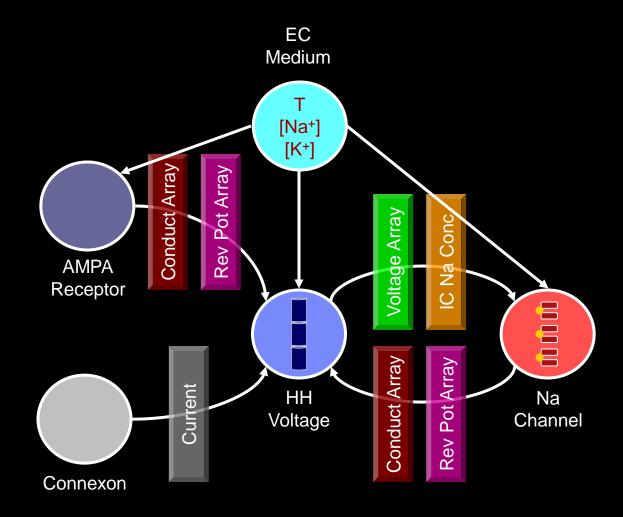
COMPARTMENT_VARIABLE_COSTS 2

Voltage 1.0

Calcium 0.95



Graph View: Synapses and Channels





Graph Specification: Channels

CHANNEL TARGETS 4

BRANCHTYPE

- O Na [Voltage] [Voltage], KDR [Voltage] [Voltage], Cah [Voltage] [Voltage, Calcium], KCa [Calcium] [Voltage]

 1 Na [Voltage] [Voltage], KDR [Voltage] [Voltage]
- 2 Cah [Voltage] [Voltage, Calcium], KCa [Calcium] [Voltage] 3 Cah [Voltage] [Voltage, Calcium], KCa [Calcium] [Voltage]

CHANNEL COSTS 4

Na 0.414243 KDR 0.254051 Cah 0.414243

KCa 0.359252

CHANNEL PARAMS 2

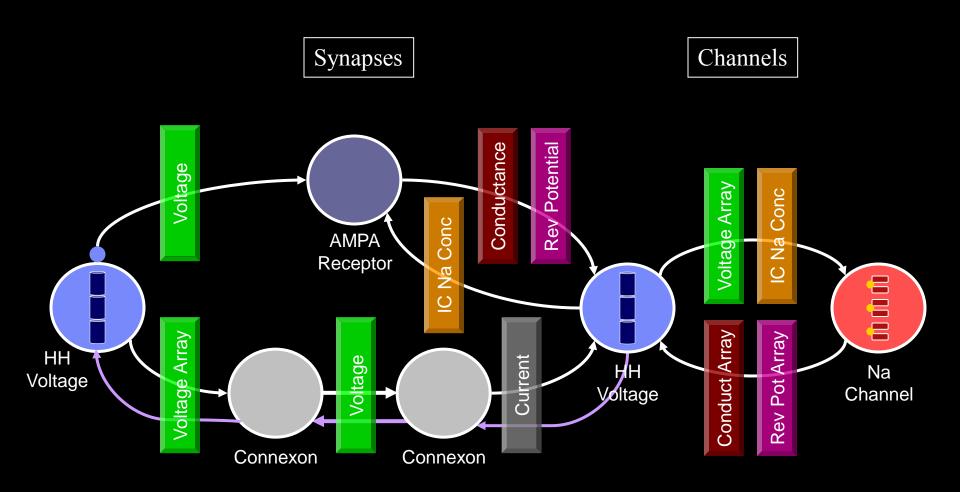
Na 2

BRANCHTYPE

 $0 < gbar = \{0.7\} >$ 1 <gbar={1.8}>



Graph View: Synapses and Channels





Graph Specification: Synapses

ELECTRICAL_SYNAPSE_TARGETS 2 **BRANCHTYPE ETYPE BRANCHTYPE ETYPE** 1 0 1 0 AxAxGap [Voltage] 0.001 2 1 2 1 DenDenGap [Voltage] 0.001

ELECTRICAL SYNAPSE COSTS 2

AxAxGap 0.005309 DenDenGap 0.005309

CHEMICAL SYNAPSE TARGETS 6

BRANCHTYPE ETYPE BRANCHTYPE ETYPE

1 1 2 0 GABAA [Voltage] [Voltage] 0.1667

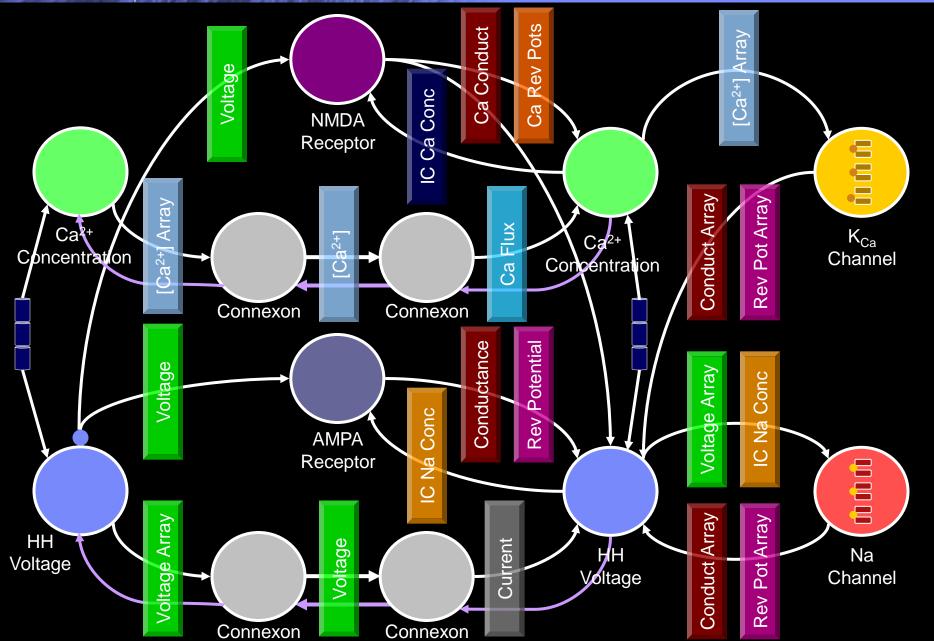
1 1 2 1 GABAA [Voltage] [Voltage] 0.1667 1 1 3 0 GABAA [Voltage] [Voltage] 0.1667 1 0 2 0 AMPA [Voltage] [Voltage] 1.0 1 0 2 1 AMPA [Voltage] [Voltage] 1.0

1 0 3 0 AMPA [Voltage] [Voltage] 1.0 NMDA [Voltage] [Voltage, Calcium] 1.0

CHEMICAL SYNAPSE COSTS 2

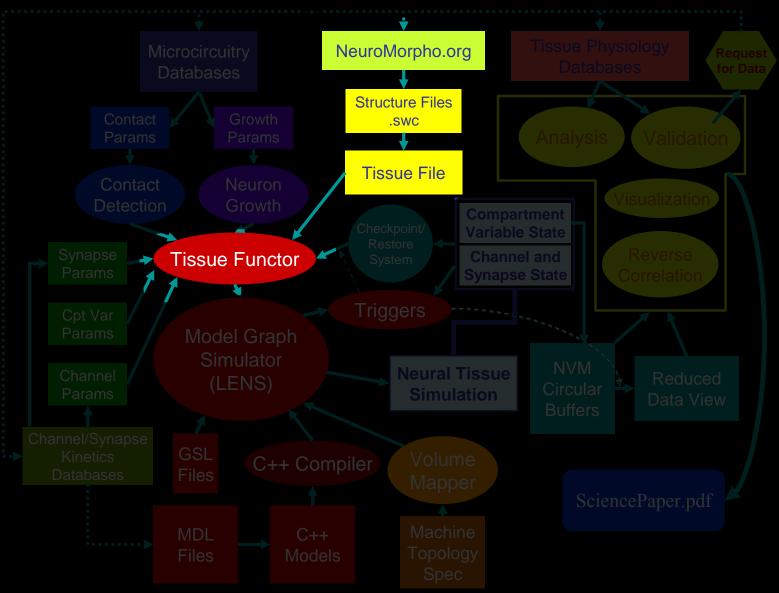
AMPA 0.296407 GABAA 0.149978



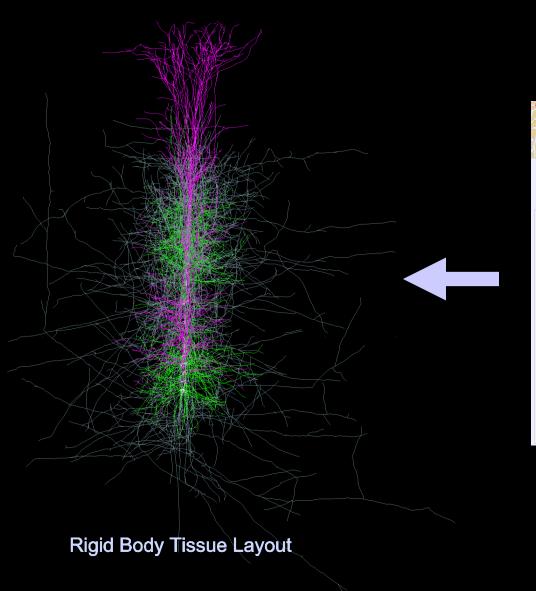


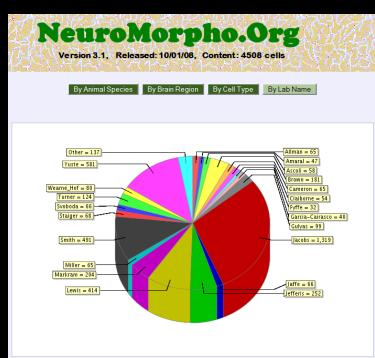


Neural Tissue Simulation Workflow



SIMULATED "MINICOLUMN"

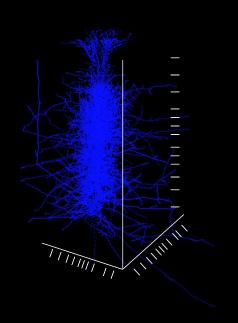




NEURAL TISSUE SIMULATION ON BLUE GENE

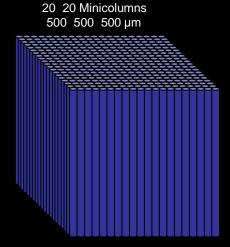
SIMULATION APPROACH:

- Distribute tissue points weighted by computational complexity
- Scale out tissue simulation across all three dimensions
- Maintain realistic neuron and synapse densities at each scale



Simulation Element	Number	Processor Balance
Neurons	1,024,000,	N/A
Branches	344,474,059	84,100 7,406
Junctions	208,947,659	51,012 4,026
Compartments	1,083,289,600	264,475 7,582
Na Channels	330,613,914	80,716 7,440
KDR Channels	330,613,914	80,716 7,440
AMPA Synapses	8,186,972,360	1,998,772 720,155
GABAA Synapses	2,255,068,948	550,553 169,064
Connexons	7,626,124	1,861 820

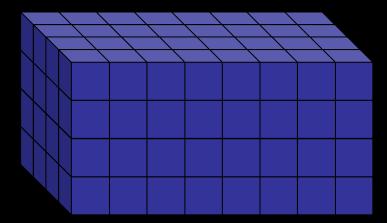
Minicolumn 20 Neurons 25 25 500 µm



Column

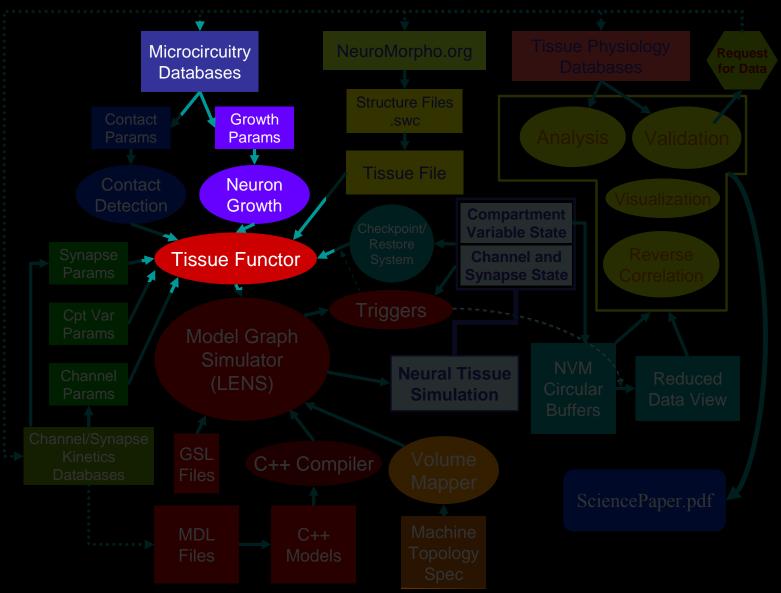
8,000 Neurons

Tissue 1,024,000 Neurons 8 4 4 Columns 4 2 2 mm





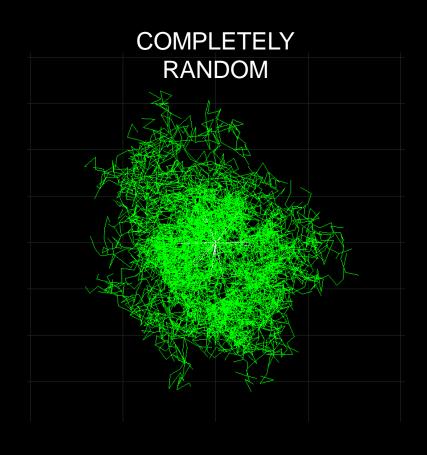
Neural Tissue Simulation Workflow



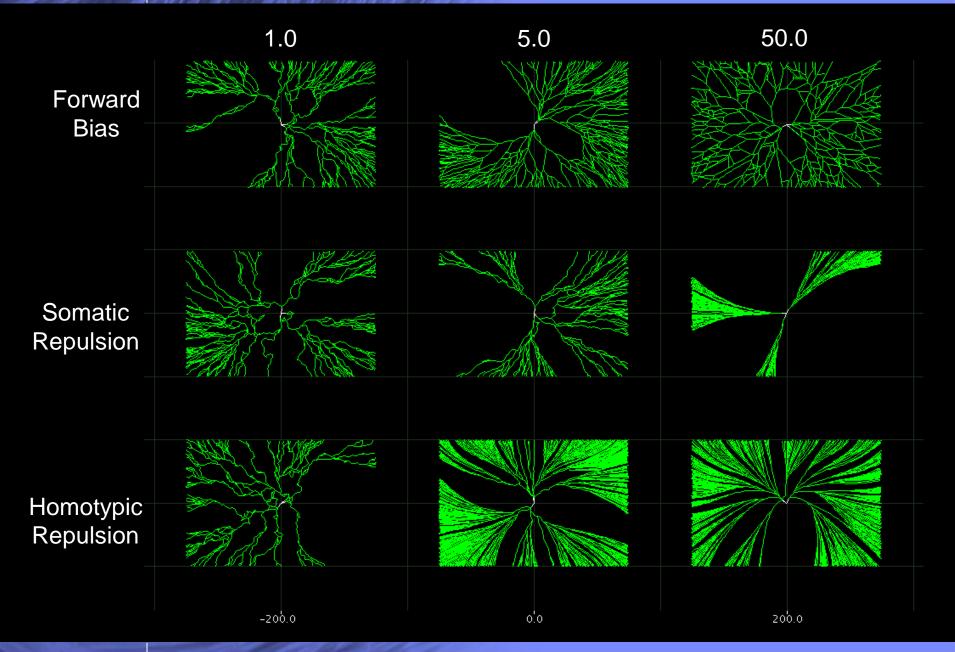


Modified Diffusion Model

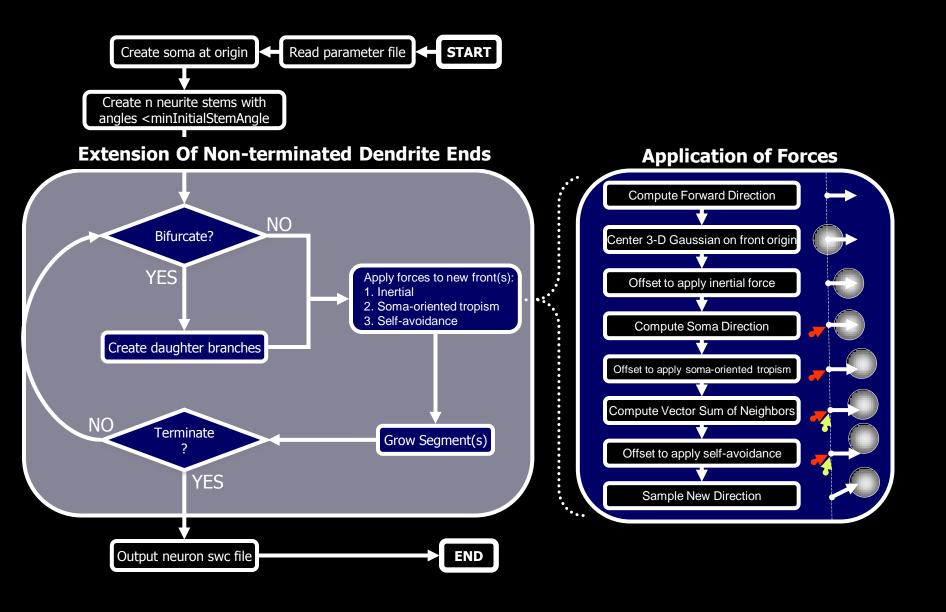
- Points sampled from 3-D
 Gaussian distribution centered on previous point
- Branching is represented as particle division
- Termination is represented by collision between particles and past trajectories as annihilation



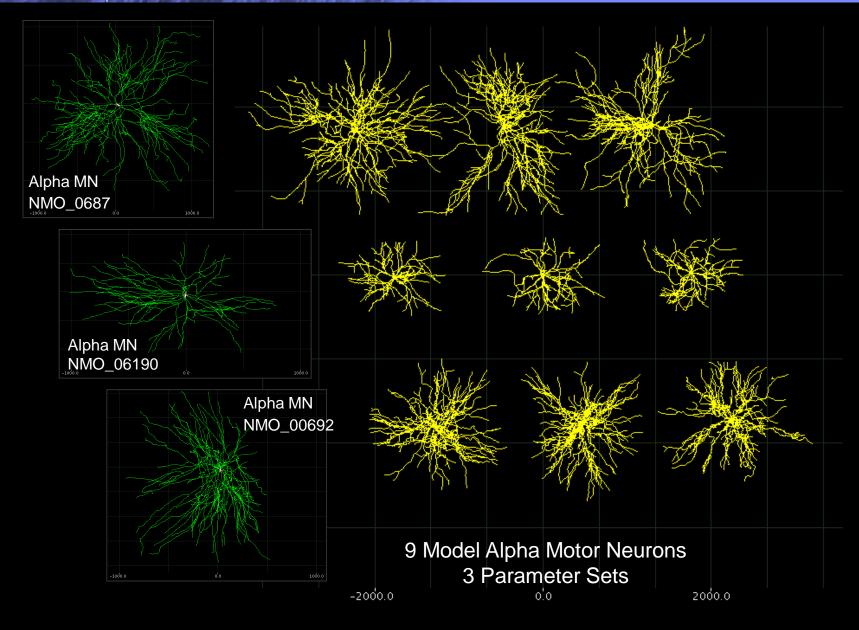




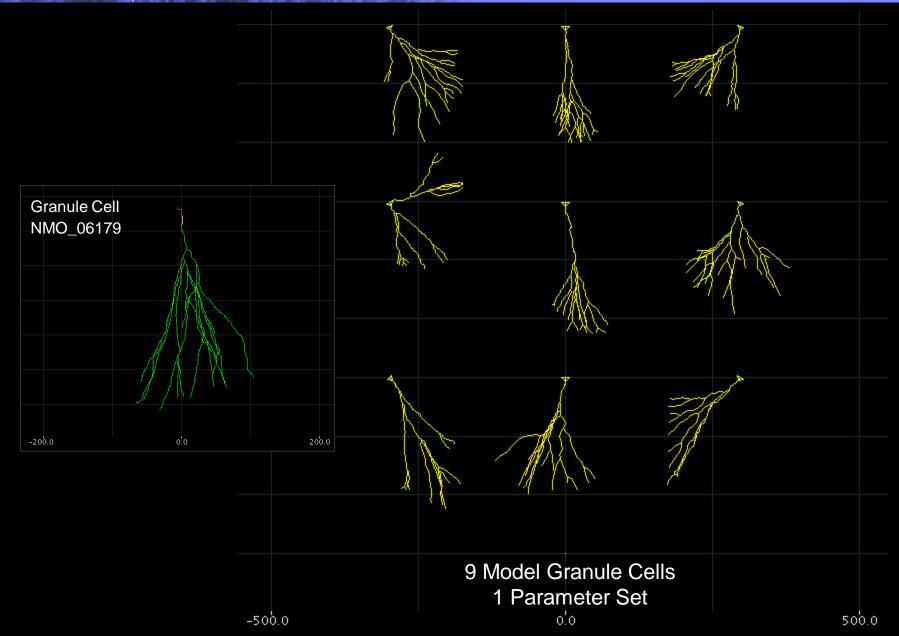


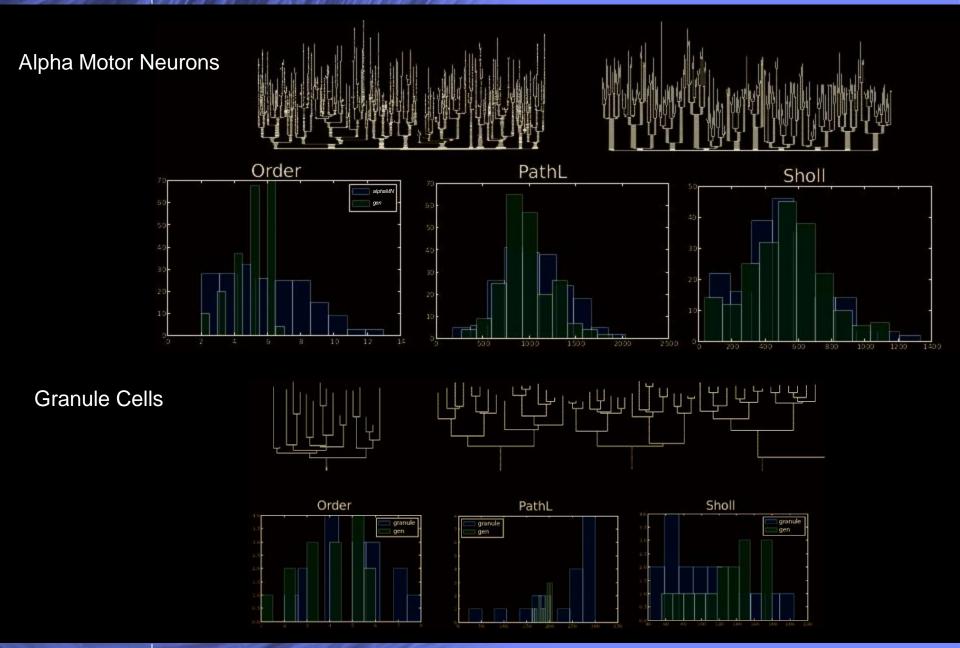




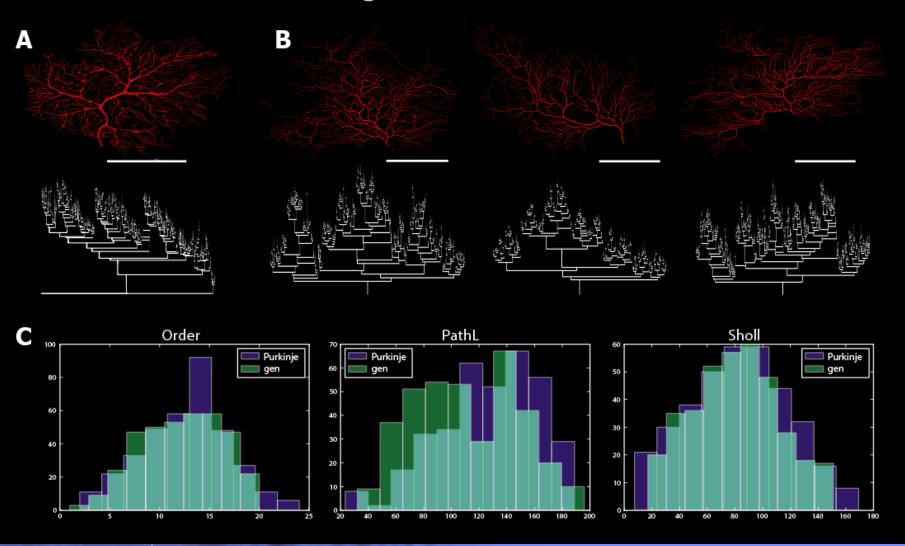






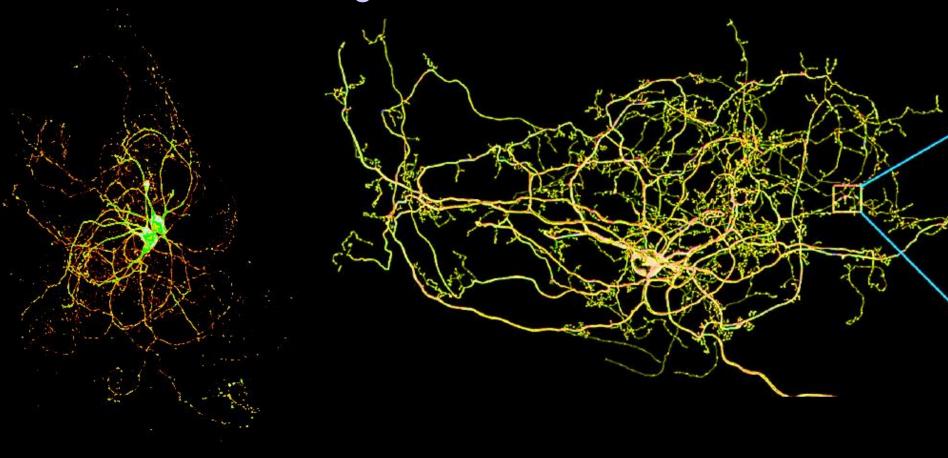


Cerebellum Modeling: Cortex



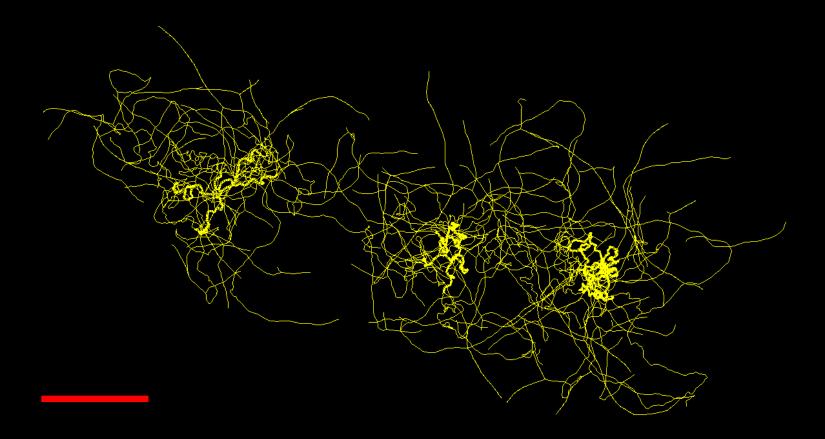


Cerebellum Modeling: Inferior Olive



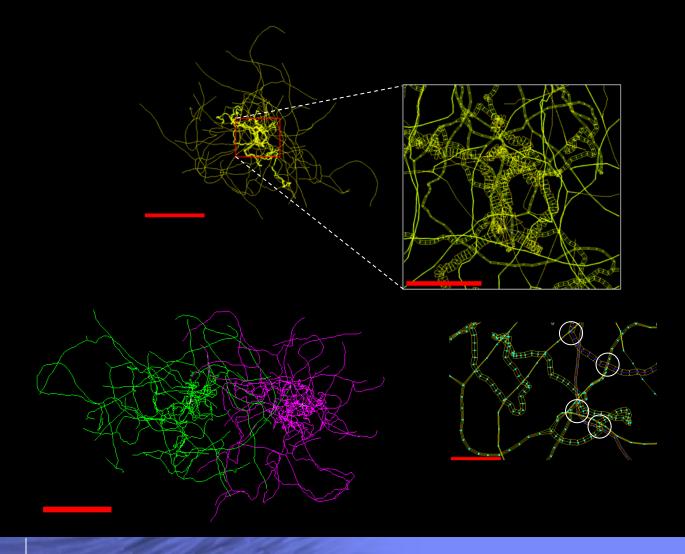


Cerebellum Modeling: Inferior Olive



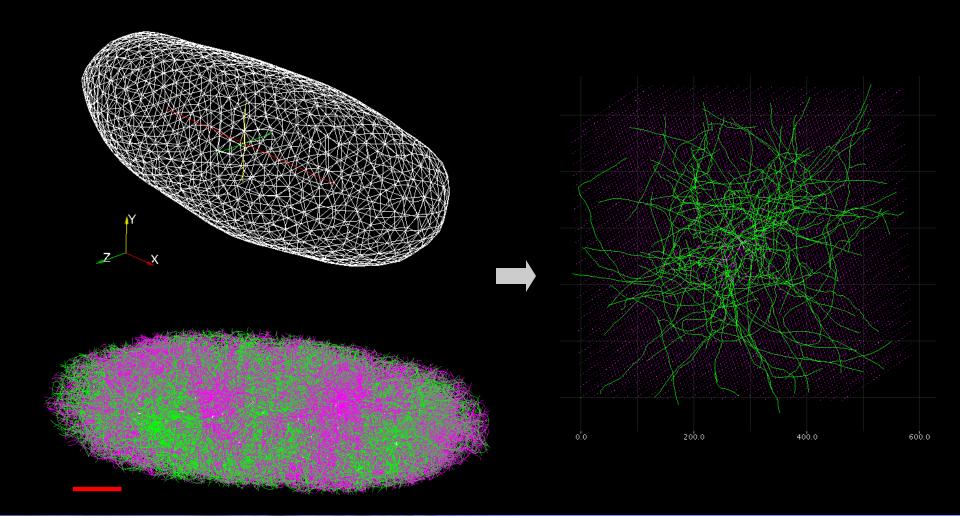


Cerebellum Modeling: Inferior Olive



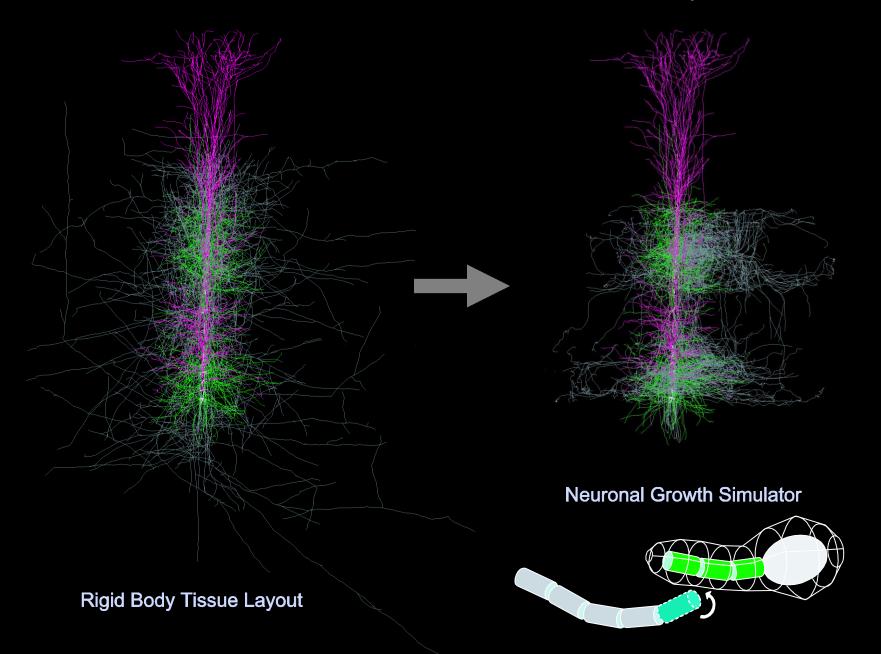


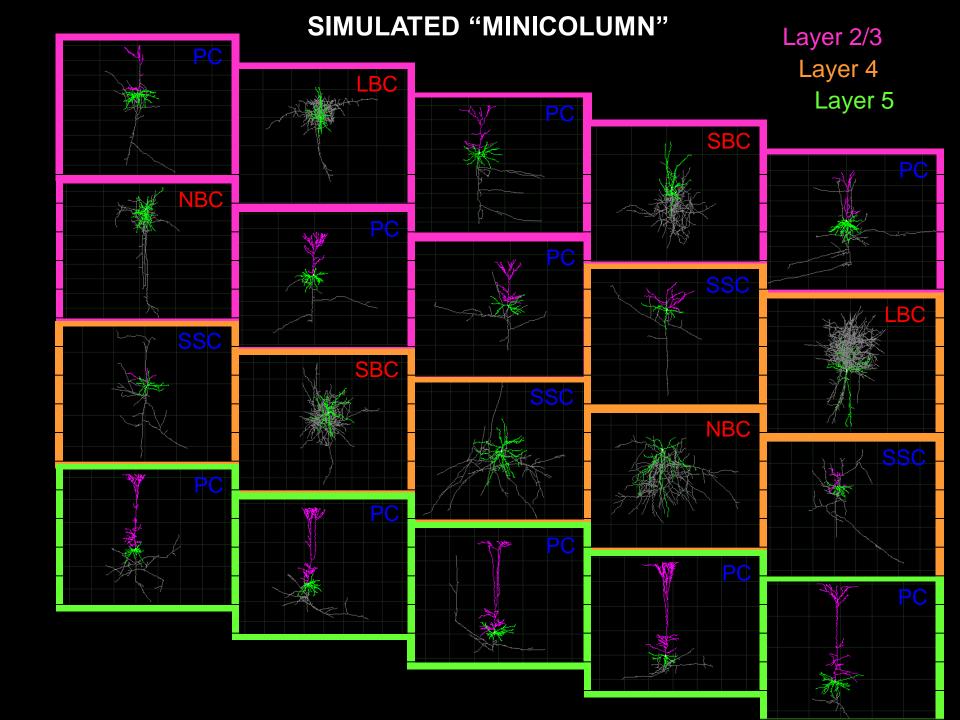
Cerebellum Modeling: Inferior Olive

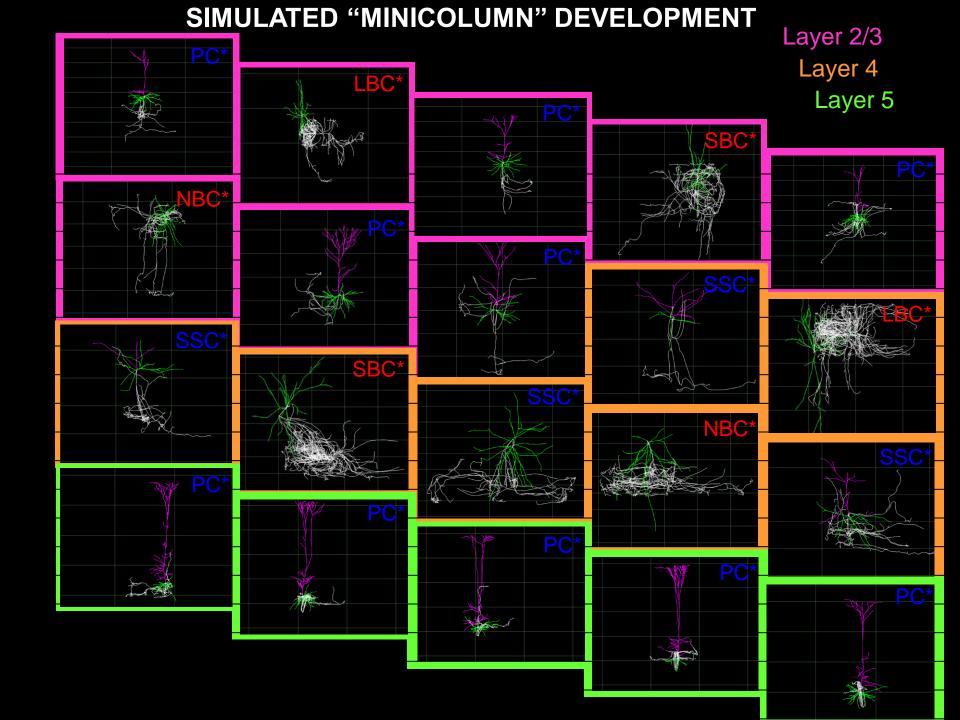


SIMULATED "MINICOLUMN" DEVELOPMENT

In collaboration with Mike Pitman, Protein Science & Molecular Dynamics









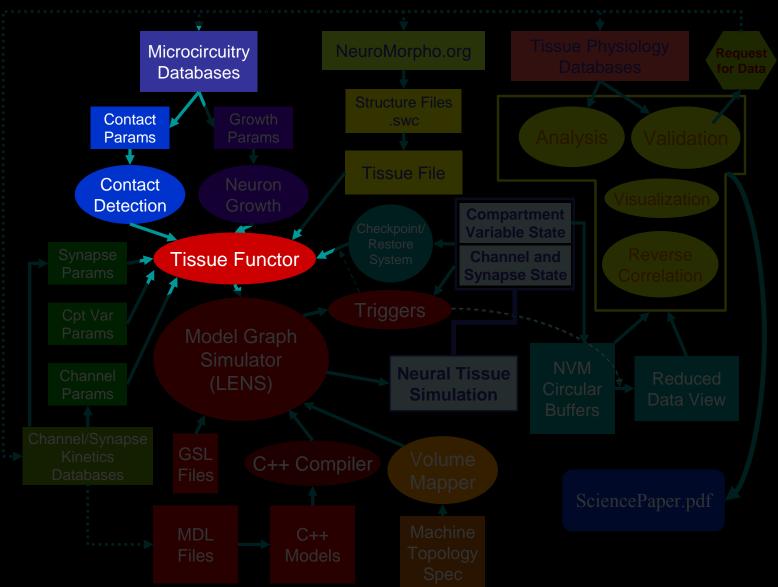
"Patient-specific models of deep brain stimulation: Influence of field model complexity on neural activation predictions A. Chaturvedi, C. R. Butson, S. F. Lempka, S. E. Cooper, C. C. McIntyre,

Brain Stimulatio Neural Tissue Simulator Approach

- Develop axonal tracts differentially using Neuron Growth Simulator, constrained by patient data
- Establish extracellular mesh of finite elements using new models expressed in MDL
- Connect finite elements to neural tissue elements using GSL
- Model effects of DBS on axonal tracts, generating action potentials in multi-compartment axonal models



Neural Tissue Simulation Workflow





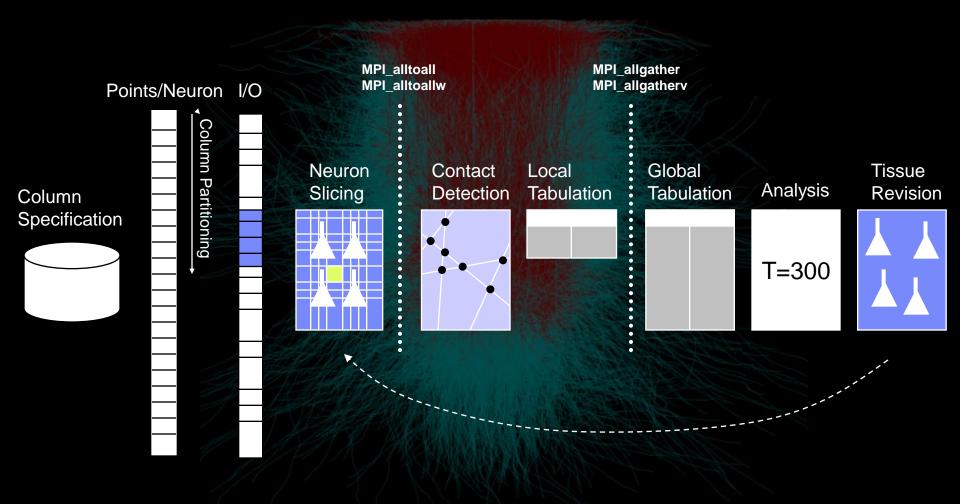
Contact Detection: The Problem

- 10,000 neurons
- 40M segments
- Average segment radius: 0.23 0.24 µm
- Average segment length: 3.65 3.2 µm
- Column dimensions: 550 1,200 550 µm
- Neurite extent: 4,400 3,300 4,300 µm
- Neuronal complexity: ≤19,000 segments
- Target compute time: ≤1 minute

Synapse creation as a function of fiber proximity: Contact Detection



Contact Detection: Architectural Overview

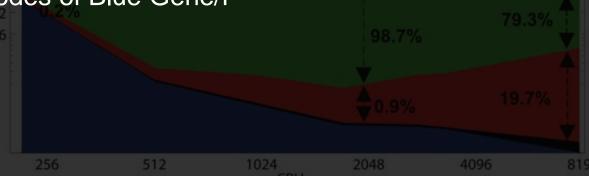




Contact Detection



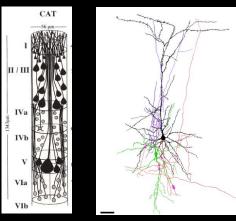
- Algorithm optimized to run multithreaded on Blue Gene/P's
- 25.5 billion contacts in 2.5 hours
- 4,096 nodes of Blue Gene/P

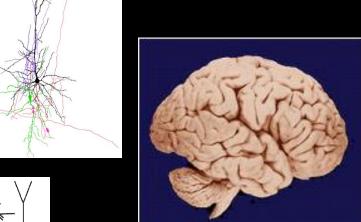


Simulation Workflow In a Single Executable

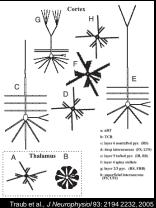
- GSL parser initializes tissue with specified grid dimensions
- Volume mapper assigns volumes to nodes of Blue Gene/P
- Tissue Functor reads tissue spec, computes consistent scheme for dividing neuron work
- On each node, Tissue Functor loads different neurons from .swc files
- Tissue Functor resamples neurons
- Communication enumerates all tissue points, constructs global point histogram, equalizes
- Neuron segments communicated to all volumes they traverse, according to histogram
- Tissue Functor executes neuron growth algorithm (optional)
- Tissue Functor executes touch detection algorithm
- Tissue Functor aggregates computational costs associated with each neuron segment
- Global cost histogram equalized in three dimensions to create a second volume slicing
- Tissue Functor communicates touches, segments to nodes responsible for models or proxies
- GSL parser creates all tissue models locally, including branches, channels, and synapses
- Probability for creating synapse models of a specific type from a set of valid touches applied
- GSL parser initializes other models such as stimulation and recording electrodes
- GSL parser interprets the specified phase structure of the simulation
- GSL parser initiates the simulation, in which each iteration comprises a sequence of phases:
 - Solve ion channel and synapse states
 - Predict branch junction states
 - Forward eliminate branches of appropriate compute orders
 - Back-substitute branches of appropriate compute orders
 - Correct branch junction states
- Between phases, state is marshalled, communicated (MPI_alltoallv), demarshalled into proxies
- Simulation terminates when the end criterion is satisfied
- All models and simulation data are destructed on all nodes of Blue Gene/P

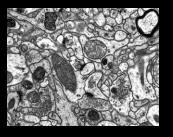






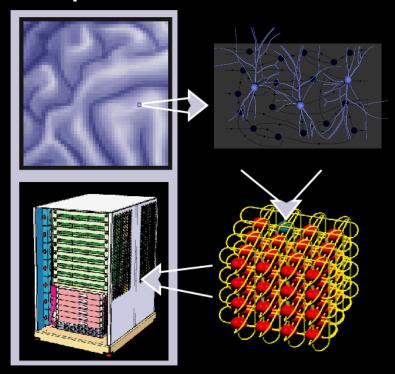








The question of axons...

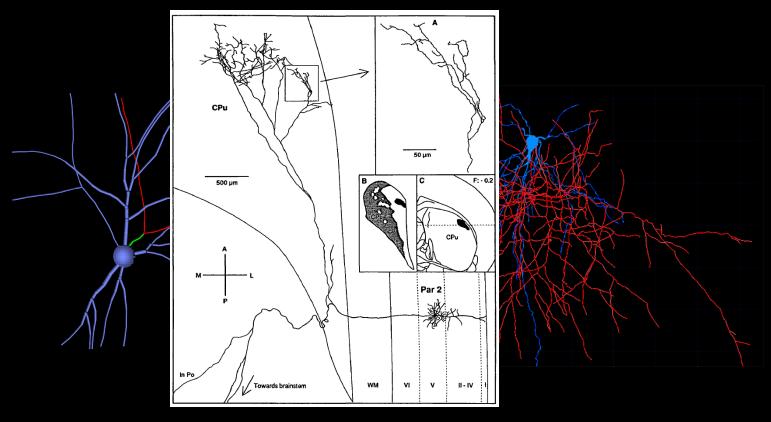


"...processors act like neurons and connections between processors act as axons..."

H. Markram. The Blue Brain Project. Nat Rev Neurosci, 7(2):153–160, Feb 2006.



The question of axons...



M. Lévesque, S. Gagnon, A. Parent, and M. Deschênes, "Axonal Arborizations of Corticostriatal and Corticothalamic Fibers Arising from the Second Somatosensory Area in the Rat

The question of axons...

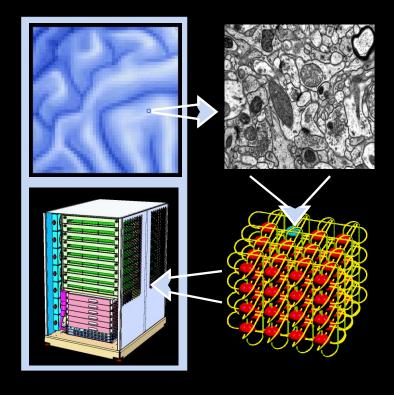
- Failures of action potential propagation can occur at certain points along an axon, introducing uncertainty surrounding the signaling role of action potentials transmitted through otherwise reliable axons [1]
- Electrical synapses between axons can initiate action potentials without first depolarizing the axon initial segment [2]
- Action potentials may be generated by a mechanism that depends on the length of the axon (e.g., bursts of action potentials of a particular duration may be generated when a calcium spike from the cell body depolarizes an axon of a particular length [1])

[1] A. Mathy, S. S. N. Ho, J. T. Davie, I. C. Duguid, B. A. Clark, and M. Husser. Encoding of oscillations by axonal bursts in inferior olive neurons. *Neuron*, 62(3):388–399, May 2009.

[2] D. Schmitz, S. Schuchmann, A. Fisahn, A. Draguhn, E. H. Buhl, E. Petrasch-Parwez, R. Dermietzel, U. Heinemann, and R. D. Traub. Axo-axonal coupling. a novel mechanism for ultrafast neuronal communication. *Neuron*, 31(5):831–840, Sep 2001.

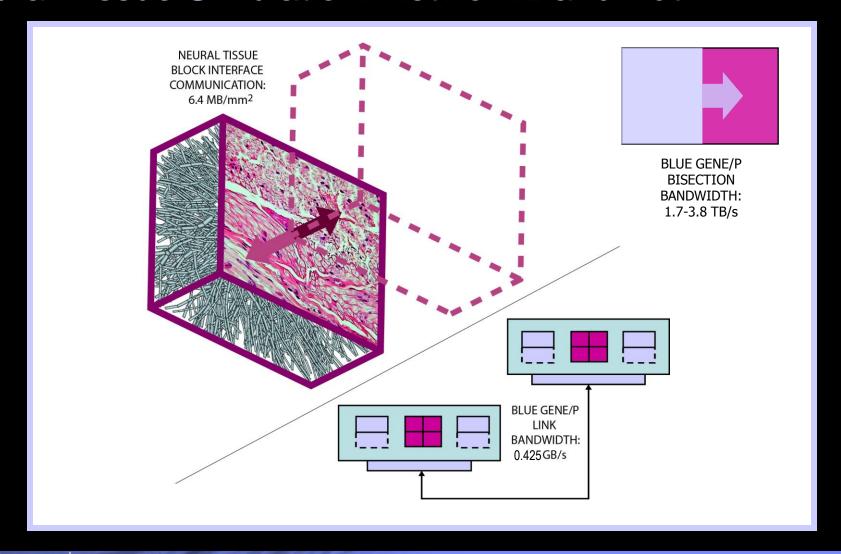
January 10, 2013







Neural Tissue Simulation: Network Bandwidth



January 10, 2013



frontiers in **NEUROINFORMATICS**

METHODS ARTICLE





An ultrascalable solution to large-scale neural tissue simulation

James Kozloski¹* and John Wagner²

- ¹ Biometaphorical Computing, Computational Biology Center, IBM Research Division, IBMT. J. Watson Research Center, Yorktown Heights, NY, USA
- ² Victorian Life Sciences Computation Initiative, IBM Research Collaboratory for Life Sciences Melbourne, Carlton, VIC, Australia

Edited by:

Markus Diesmann, RIKEN Brain Science Institute, Japan

Reviewed by:

Michael Hines, Yale University, USA Abigail Morrison, Bernstein Center Freiburg, Germany

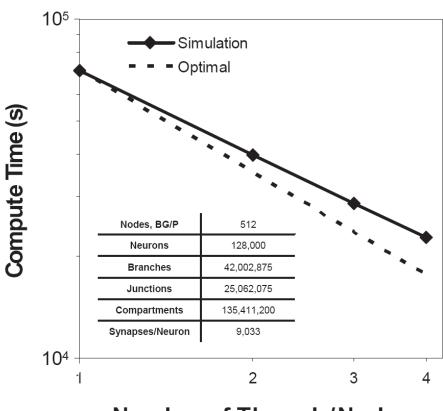
*Correspondence:

James Kozloski, Biometaphorical Computing, Computational Biology Center, IBM Research Division, IBM T. J. Watson Research Center, 1101 Kitchawan Road, Room 05-144, Yo Heights, NY, USA. e-mail: kozloski@us.ibm.com Neural tissue simulation extends requirements and constraints of previous neuronal and neural circuit simulation methods, creating a tissue coordinate system. We have developed a novel tissue volume decomposition, and a hybrid branched cable equation solver. The decomposition divides the simulation into regular tissue blocks and distributes them on a parallel multithreaded machine. The solver computes neurons that have been divided arbitrarily across blocks. We demonstrate thread, strong, and weak scaling of our approach on a machine with more than 4000 nodes and up to four threads per node. Scaling synapses to physiological numbers had little effect on performance, since our decomposition approach generates synapses that are almost always computed locally. The largest simulation included in our scaling results comprised 1 million neurons, 1 billion compartments, and 10 billion conductance-based synapses and gap junctions. We discuss the implications of our ultrascalable Neural Tissue Simulator, and with our results estimate requirements for a simulation at the scale of a human brain.

Keywords: neural tissue, simulation, parallel computing, distributed computing, Hodgkin–Huxley, numerical methods, ultrascalable, whole-brain

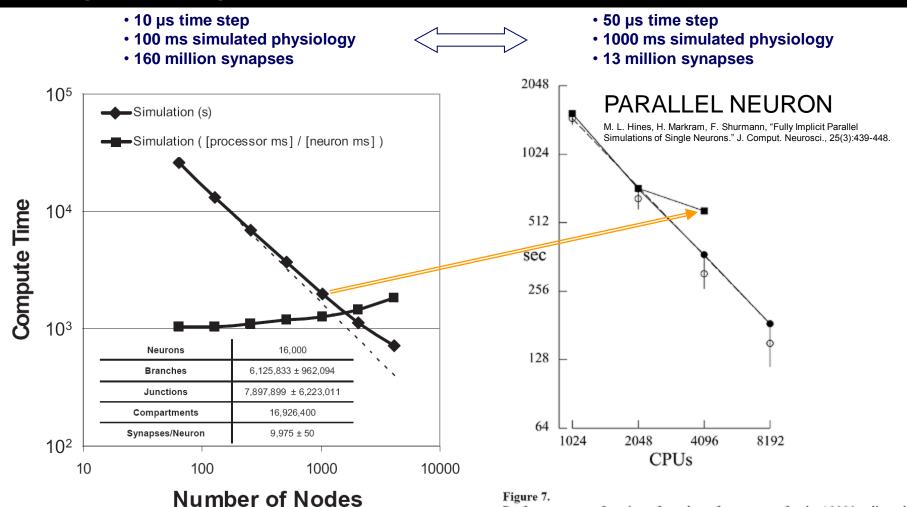


Thread Scaling



Number of Threads/Node

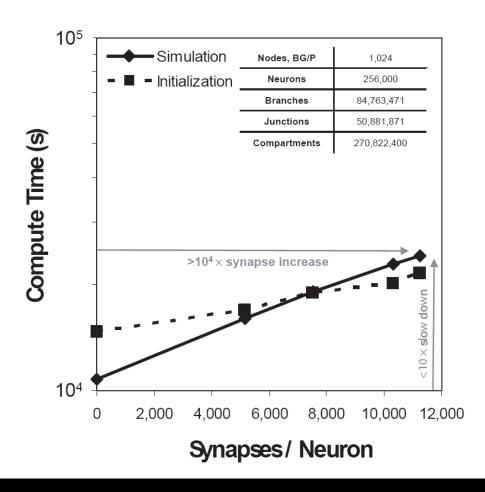
Strong Scaling



Performance as a function of number of processors for the 10000 cell model.

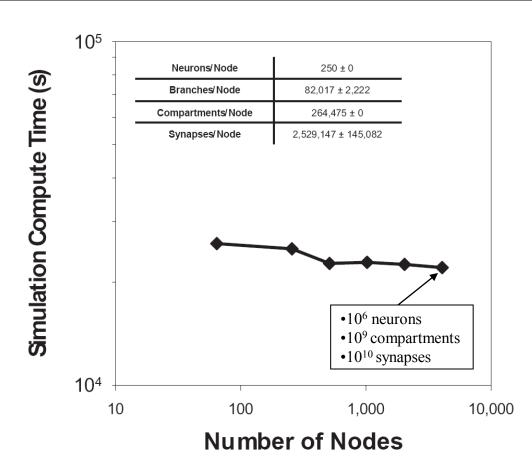


Synapse Scaling



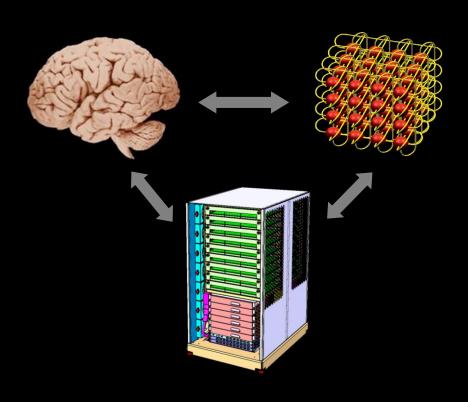


Weak Scaling





Feasibility of Human Brain-Scale Calculations

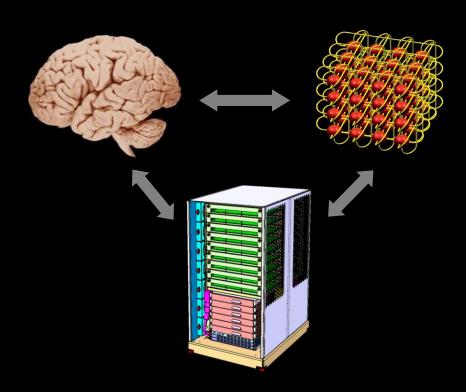


1 Liter of Neural Tissue In 10⁷ Volumes

- 10⁸ µm³ / volume
- 10⁶ compartments / volume, assuming 100 µm³ / compartment
- 10¹⁰ flop / 50 µs simulation timestep / volume, assuming 10⁴ flop / timestep / compartment (includes Hodgkin Huxley cable solution, plus 10 ion channels or synapses / compartment)
- 64kB communicated / volume face / timestep, assuming 32 bytes communicated / spanning compartment in each direction, and 10 µm² / compartment cross section
- 2.25 GB of memory / volume, including 250 MB of simulation overhead, plus 1.60 kB / compartment and 64 bytes / channel or synapse



Feasibility of Human Brain-Scale Calculations



10⁷ volumes mapped to a hypothetical Blue Gene with 10⁷ computational nodes

- 109 flops / node (Blue Gene/P scale)
- 6.4 kB/s of link bandwidth (in each direction, to accommodate packet overhead, well within Blue Gene/P scale)
- 3 GB of memory / node (Blue Gene/P scale);
 30 PB of total machine memory
- 6 GB/s memory bandwith, assuming all simulation state must be traversed 3 times in each simulation time step (Blue Gene/P scale)
- Simulation time to real time factor of 2 10⁵ yielding a simulated time duration of 400 ms per day of computation, or 3 s per week of computation



"Brain Systems Computation": Approach

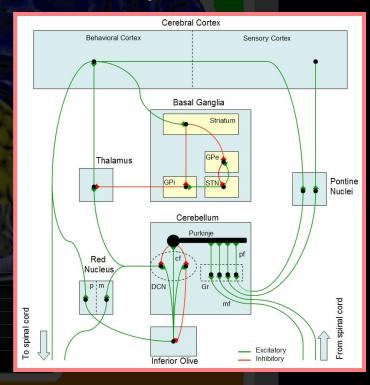
- Start with whole brain structures at the scale of BG/P
- Add brain system components with other whole brain structures
- Analyze and validate based on system level observations
- Scale system as resources grow towards exascale





"Brain Systems Computation": Example

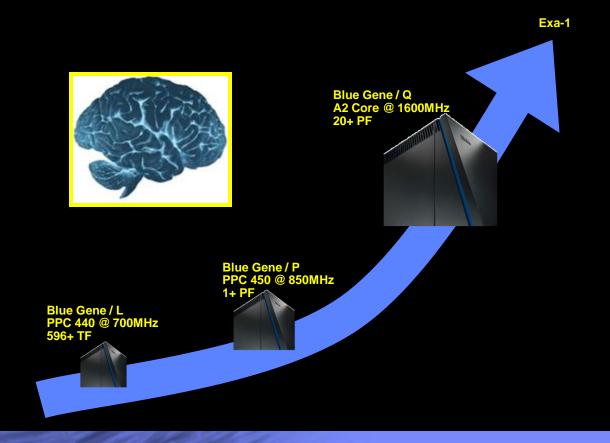
- Inferior Olive
- Deep Cerebellar Nuclei
- Cerebellar Cortex
- Thalamocortical
- Pontine Nuclei
- Rubrospinal
- Basal Ganglia





Mapping To Exascale

	GFLOPS	GFLOPS (App)	mem/node (GB)	LINK BW (GB/s)	mem BW (GB/s)
BG/L	5.6	1.12	0.5	0.175	5.6
BG/P	13.6	2.72	4	0.425	13.6
BG/Q	204.8	40.96	16	2.0	43
Exa-1 (Required)	1000	200	16	6.8	300



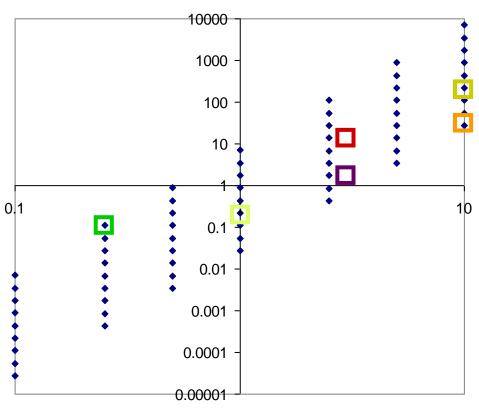


Speed Requirements By Tissue Size

APPLICATION REQUIREMENTS



GFLOPS / node



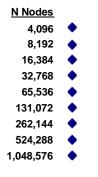
MACHINE PROPERTIES

- 0.02 mL, 4 racks BG/P
 - 1 mL, 72 racks BG/P
- 27 mL, 288 racks BG/P with 15 GB NVM/node
- 27 mL, 28 racks BG/Q with 150 GB NVM/node
- 1 L, 1,000 racks BG/Q with 150 GB NVM/node
 - 1 L, 144 racks Exa-1 with 1 TB NVM/node

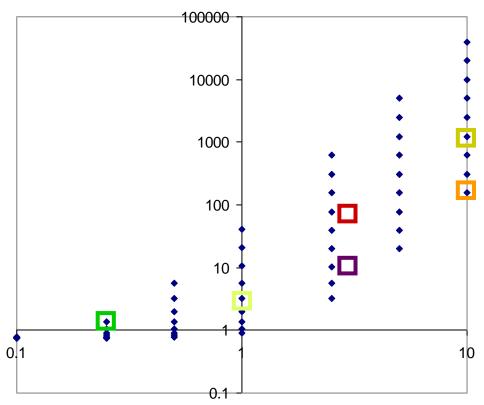


Memory Requirements By Tissue Size (mL⁻³)

APPLICATION REQUIREMENTS



Memory / node (GB)

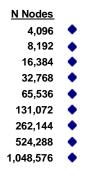


MACHINE PROPERTIES

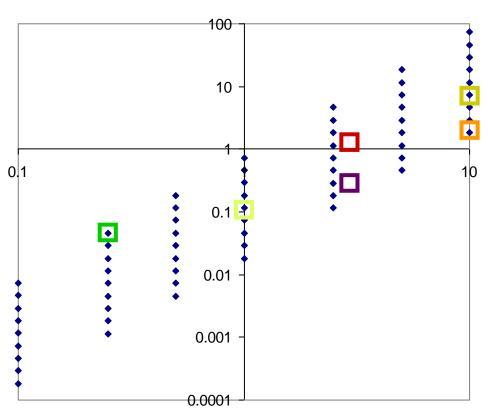
- 0.02 mL, 4 racks BG/P
- 1 mL, 72 racks BG/P
- 27 mL, 288 racks BG/P with 15 GB NVM/node
- 27 mL, 28 racks BG/Q with 150 GB NVM/node
- 1 L, 1,000 racks BG/Q with 150 GB NVM/node
 - 1 L, 144 racks Exa-1 with 1 TB NVM/node

Link BW Requirements By Tissue Size (mL⁻³)





Link BW (GB/s)



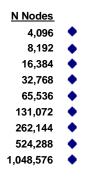
MACHINE PROPERTIES

- 0.02 mL, 4 racks BG/P
 - 1 mL, 72 racks BG/P
- 27 mL, 288 racks BG/P with 15 GB NVM/node
 - IIIL, 200 Tacks BG/P With 15 GB NVM/hode
- 27 mL, 28 racks BG/Q with 150 GB NVM/node
- 1 L, 1,000 racks BG/Q with 150 GB NVM/node
 - 1 L, 144 racks Exa-1 with 1 TB NVM/node

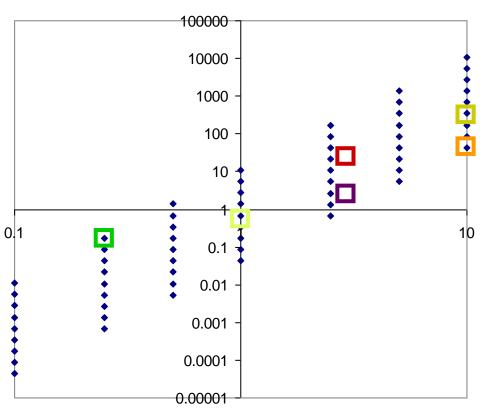


Memory BW Requirements By Tissue Size (mL⁻³)





Mem BW (GB/s)

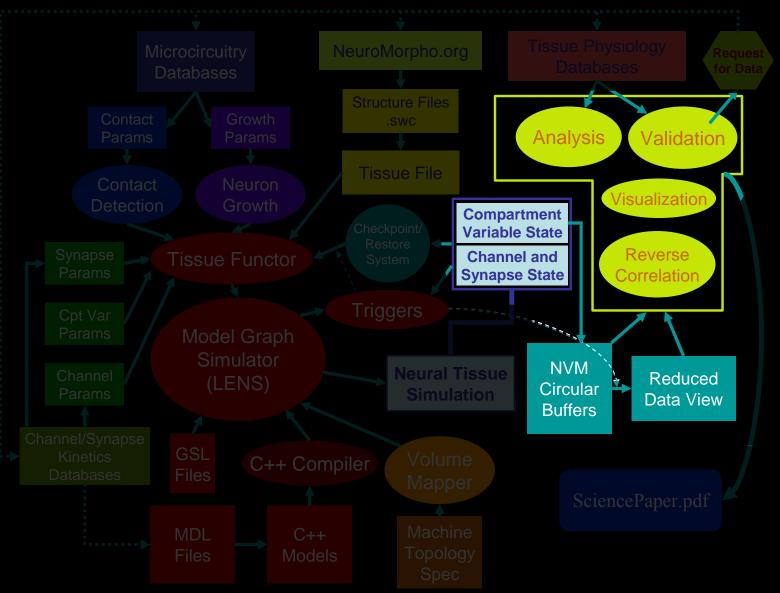


MACHINE PROPERTIES

- 0.02 mL, 4 racks BG/P
 - 1 mL, 72 racks BG/P
- 27 mL, 288 racks BG/P with 15 GB NVM/node
- 27 mL, 28 racks BG/Q with 150 GB NVM/node
- 1 L, 1,000 racks BG/Q with 150 GB NVM/node
 - 1 L, 144 racks Exa-1 with 1 TB NVM/node



Neural Tissue Simulation Workflow



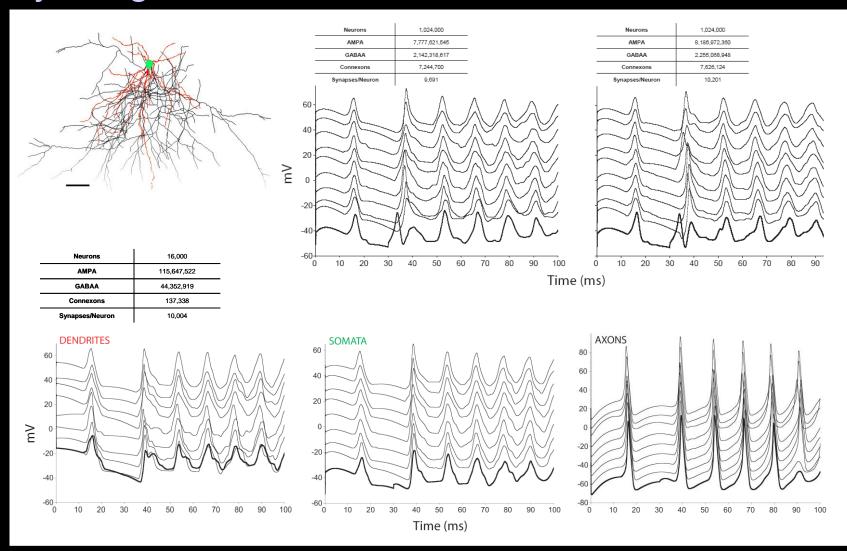


Neural Tissue Simulator: Current Storage Model

- Model Graph Simulator creates special data collection graph elements, specified in MDL, declared in GSL
- Data collectors' connections to specific neural tissue elements declared in GSL according to key specification
- At runtime, the existence of elements on each compute node matching key is established, and data collectors created
- Data collectors write to own files upon satisfaction of predicates of GSL-specified triggers (e.g., iteration)

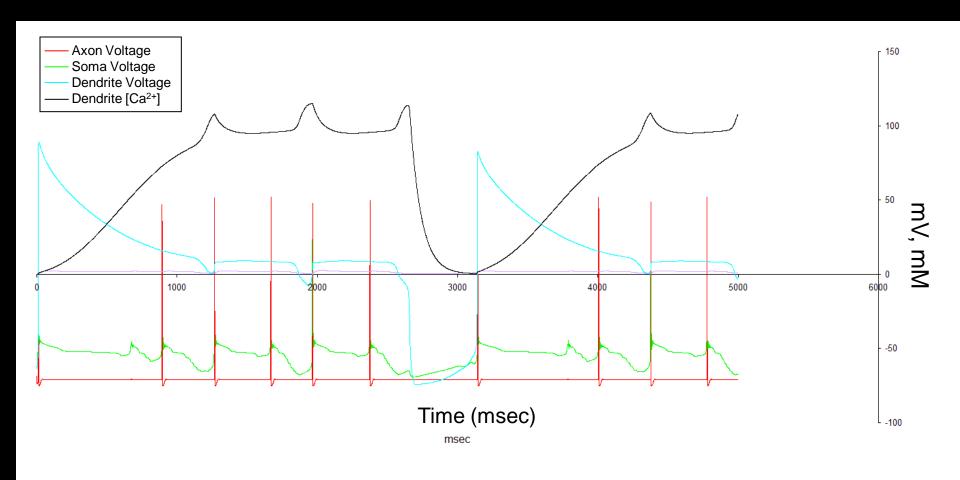


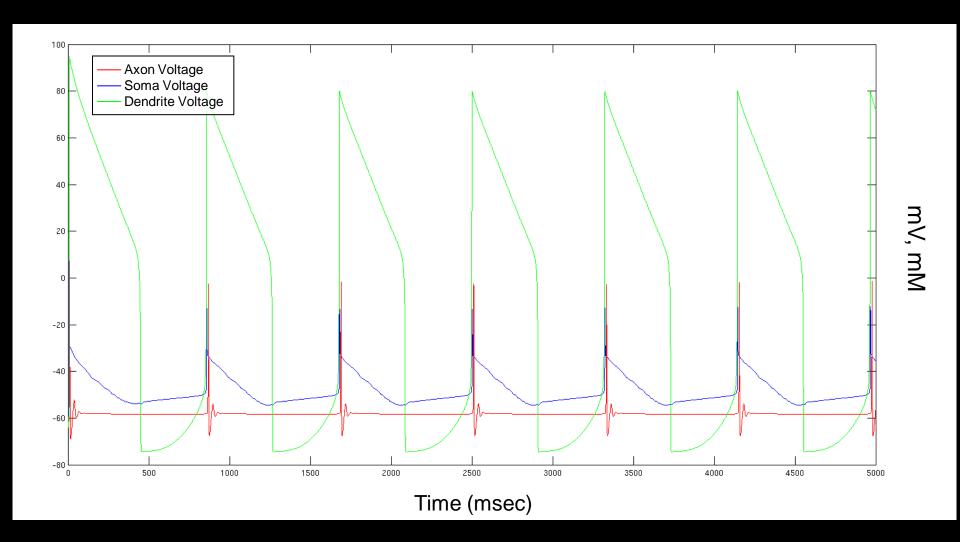
Physiological Results: "Neocortical" Simulation





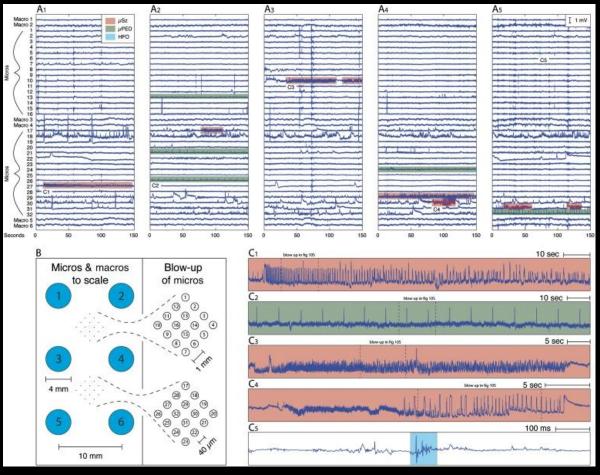
Physiological Results: Inferior Olive Simulation







Neural Current Analyzer: I/O Bound Application



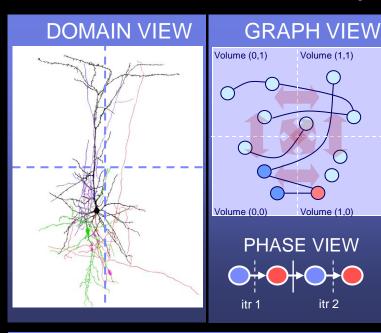
Microseizures and the spatiotemporal scales of human partial epilepsy

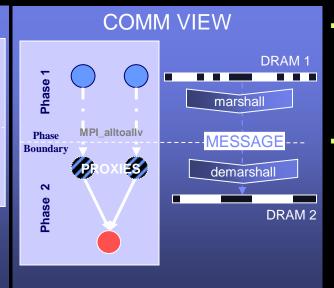
Matt Stead, Mark Bower, Benjamin H. Brinkmann, Kendall Lee, W. Richard Marsh, Fredric B. Meyer, Brian Litt, Jamie Van Gompel, and Greg A. Worrell

- Standard neural analysis methods lack the ability to store massive data quickly, perform postsimulation analyses to discover causation
- Standard demonstrationdriven approaches record exemplars to test hypotheses inherent in simulation design
- A discovery-driven approach and data analysis framework would allow the quantitative identification of causes of network phenomena in neural tissue

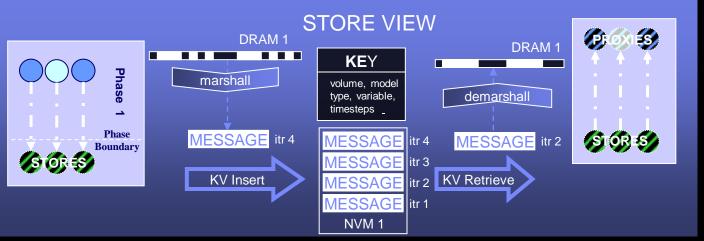


Neural Current Analyzer: Proposed Storage Model





- Simulation framework creates messages at phase boundaries to be sent across node boundaries to initialize proxies for next phase
- Storage model uses framework generated messages to store state for
 - –Checkpoint rollback
 - -Triggered analysis
 - -Causality analysis
 - -Visualization
- Framework marshalling generates key/value inserts saving graph-node state
- Framework uses demarshalling infrastructure to generate key/value/offset retrieve to initialize graphnode proxy state as required by analysis algorithm



January 10, 2013



Neural Current Analyzer: Application Areas

Epilepsy

- Quantitative analysis of current motifs responsible for epileptogenesis
- Identification of specific mechanisms of observed channelopathies
- Drug candidate identification
- Brain stimulation protocols and parameters

Predictive EEG modeling

- Solve EEG in FEM model of extracellular fields for each current motified
- Use EEG "motifs" as building blocks (components) for a forward model of real EEG recordings
- Relate EEG to behavior to predict functional brain (current) states



Acknowledgements

- Blake Fitch, Volume Decomposition Design, Neural Current Analyzer on BGAS
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- Benjamin Torben-Nielsen (EPFL) and Heraldo Memelli (Stony Brook),
 Constrained Diffusion Model of Neuron Morphogenesis
- Mike Pitman, Molecular Dynamics abstractions of neural growth and development
- Maria Eleftheriou, Model Graph Simulator Testing