Simulating Neuronal Networks with PyNEST

It is easy to do – but do we know what we are doing?

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[simula . research laboratory]



Overview

- NEST Initiative & Simulator
 - PyNEST: Simple examples
 - PyNEST: Network examples
 - "Showtime"
- How "science" are simulations today?
 - Reproducibility
 - Sharing and re-use
 - Perspectives

NEST Initiative & Simulator

The NEST Initiative



- <u>Neural Simulation Technology (NEST)</u> Initiative was established in 2001.
- Currently four core developing groups:
 - 1. Honda Research Institute Europe
 - 2. BCCN Freiburg
 - 3. Norwegian University of Life Science, Ås
 - 4. Brain Science Institute, RIKEN

- Goals of the NEST Initiative:
 - 1. share expertise
 - 2. combine resources
 - 3. coordinate research and software development
 - 4. increase software quality
 - 5. share simulations and results
 - Activities:
 - http://www.nest-initiative.org
 - Public releases of NEST available since fall 2004
 - Since 2004 used at the EU Advanced Course in Computational Neuroscience
 - NEST Workshops at the CNS*2005 in Madison (USA) and CNS*2006 in Edinburgh (UK)

Computational Neuroscience

"The goal of neural modeling is to relate, in nervous systems, function to structure on the basis of operation."

MacGregor & Lewis 1977

Important properties

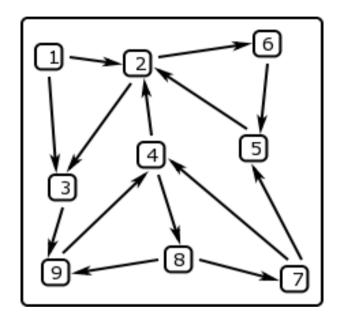
- Discrete processing units: *neurons*
- Neurons process incoming signals internally
- Neurons communicate through stereotypical pulses with finite transmission delays: *spikes*
- Typical scales
 - internal dynamics ~ 1ms
 - spike delays ~ 1ms
 - firing rates ~ 10 Hz
 - 10⁵ neurons/mm³
 - 10⁴ inputs/output per neuron

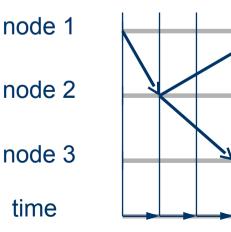
NEST in a nutshell

- Available from www.nest-initiative.org
- Command-line application
- Network models built from neurons, synapses, and devices.
- High-level simulation language (Python or SLI).
- Models for neurons, synapses, and devices are written in C++
- Support for parallel and distributed simulation (Threads and MPI)
- Used at international summer schools since 2004
- «Tell us and cite us» open source license

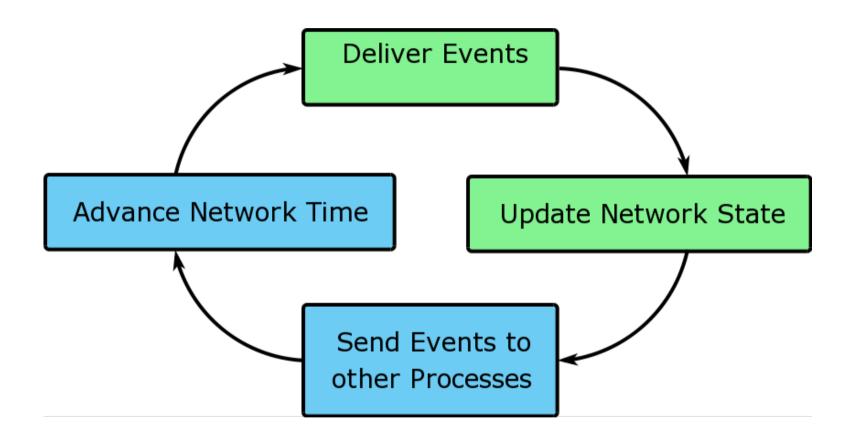
Neural Networks in NEST

- Network: Directed graph
- Nodes: Neurons (10⁴-10⁶)
- Edges: Connections (10⁷-10⁹)
- Interaction: Delayed pulses
- Kernel tasks:
 - Process incoming spikes
 - Advance neuron states
 - Emit outgoing spikes





Parallel Simulation in NEST2



SERIAL

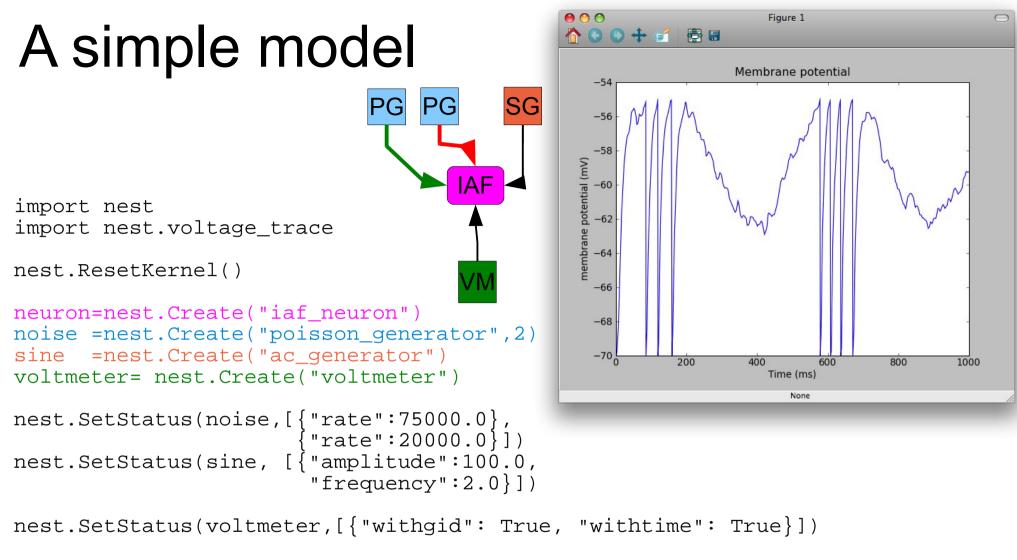


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NEST

Simple Examples



```
nest.ConvergentConnect(noise,neuron,weight=[1., -1.])
nest.Connect(voltmeter,neuron)
nest.Connect(sine,neuron)
```

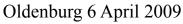
```
nest.Simulate(1000.0)
```

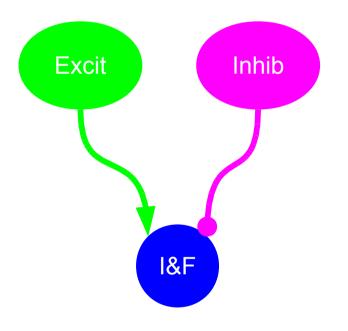
nest.voltage_trace.from_device(voltmeter)

```
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```

Example: Optimizing a network

- Excitatory population modeled as Poisson process
- Inhibitory population modeled as Poisson process
- Single I&F neuron receiving input from both populations
- Goal: Adjust inhibitory population rate so neuron fires with same rate as excitatory population
- Approach: repeated simulation
 + bisection





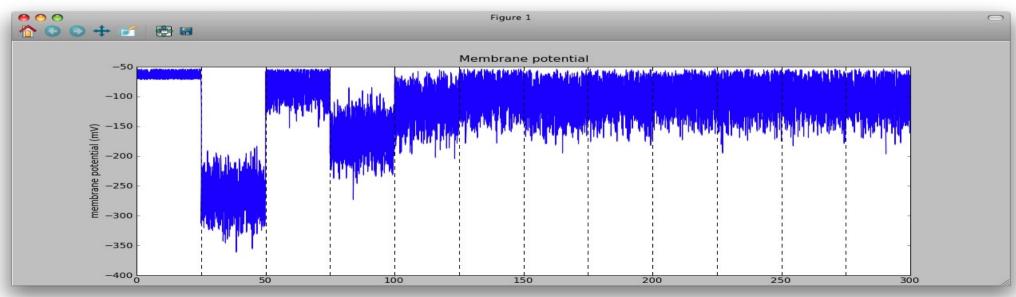
Optimizing: the code

```
= nest.Create("iaf neuron")
neuron
              = nest.Create("poisson_generator",2)
noise
spikedetector = nest.Create("spike detector")
nest.SetStatus(noise, [{"rate": n_ex*r_ex}, {"rate": n_in*r_in}])
nest.ConvergentConnect(noise, neuron, [w excit, w inhib], 1.0)
nest.Connect(neuron, spikedetector)
in rate = bisect(lambda x: output rate(x) - r ex,
                 lower, upper, xtol=prec)
def output rate(quess):
    rate = float(abs(n_in*guess))
    nest.SetStatus([noise[1]], "rate", rate)
    nest.SetStatus(spikedetector, "n events", 0)
    nest.Simulate(t sim)
    out=nest.GetStatus(spikedetector, "n events")[0]*1000.0/t sim
    return out
```

Optimization: example run

Inhibitory	rate	estimate:	15.00	Hz	->	Neuron	rate:	347.64	Hz
Inhibitory	rate	estimate:	25.00	Ηz	->	Neuron	rate:	0.04	Hz
Inhibitory	rate	estimate:	20.00	Ηz	->	Neuron	rate:	37.04	Hz
Inhibitory	rate	estimate:	22.50	Hz	->	Neuron	rate:	0.00	Hz
Inhibitory	rate	estimate:	21.25	Hz	->	Neuron	rate:	0.92	Hz
Inhibitory	rate	estimate:	20.62	Hz	->	Neuron	rate:	7.32	Hz
Inhibitory	rate	estimate:	20.94	Hz	->	Neuron	rate:	3.48	Hz
Inhibitory	rate	estimate:	20.78	Hz	->	Neuron	rate:	3.92	Hz
Inhibitory	rate	estimate:	20.70	Hz	->	Neuron	rate:	6.04	Hz
Inhibitory	rate	estimate:	20.74	Hz	->	Neuron	rate:	5.76	Hz
Inhibitory	rate	estimate:	20.76	Hz	->	Neuron	rate:	5.24	Hz
Inhibitory	rate	estimate:	20.77	Hz	->	Neuron	rate:	5.28	Hz

Optimal rate for the inhibitory population: 20.77 Hz



PyNEST

Network Example

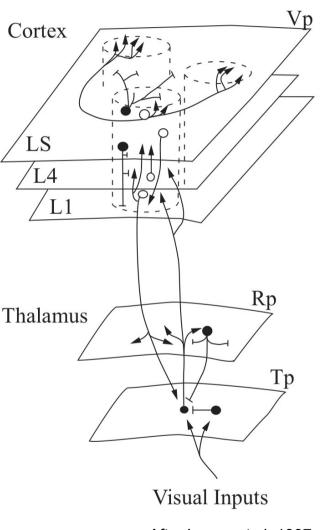
NEST Topology Module

- Idea: User-friendly support for layered networks
- Implementation: Kittel Austvoll
- Describe network as collection of layers
- Elements of a layer can be
 - Individual neurons
 - Groups of neurons (e.g. Microcolumn)
 - Placed on a fixed grid
 - Placed arbitrarily in space
- Connections described by
 - Masks: no connections outside mask
 - Kernels: give distance-dependent connection probability

Example Network

(after Lumer et al, 1997)

- LGN: Layer of individual neurons
- Vp: Layer of microcolums
 - L4: 2 pyr. cells, 1 internrn
 - L6: 1 pyramidal cell
- Connections:
 - LGN -> Vp/L4 pyr: rectangle
 - Vp/L4 pyr -> internrn: circular
 - Vp/L4 internrn -> pyr: "doughnut"
 - Vp/L4 internrn -> internrn: "flat"
 - Vp/L4 pyr -> Vp/L6 pyr: Gaussian
 - Vp/L6 pyr -> LGN: Gaussian



PyNEST/Topology Code for Network

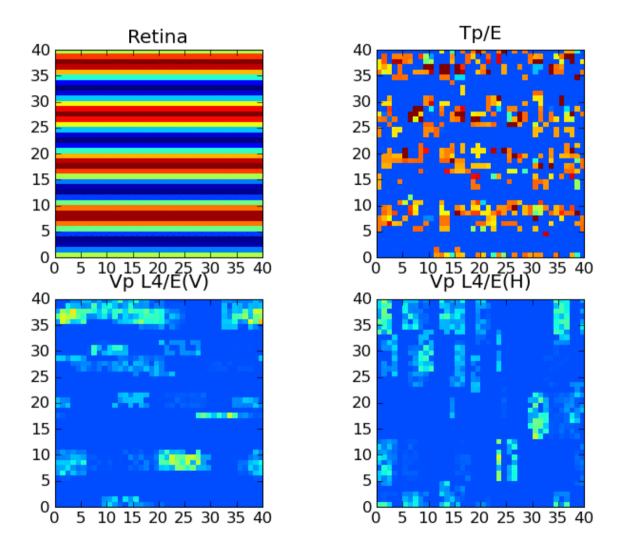
```
layout = {'rows': 50, 'columns': 50,
                                                                                                                                                                                                        # - Retina -> LGN -----
                                     'extent': [5.0, 5.0],
                                     'center': [0.0, 0.0],
                                     'edge wrap': False}
 # - Retina ------
                                                                                                                                                                                                                  'delay': 1.0,
                                                                                                                                                                                                                  'weight': 10.0})
nest.CopyModel('ac poisson generator',
                                                      'retina cell',
                                                       {'DC': 30.0, 'AC': [30.0],
                                                           'Freq': [2.0], 'Phi': [0.0]})
ret config = dict(layout)
ret config['elements'] = 'retina cell'
retina = nest.CreateLayer(ret config)
                                                                                                                                                                                                                   'kernel': 0.5,
 # - LGN -----
                                                                                                                                                                                                                  'weights': 5.0,
nest.CopyModel('iaf neuron', 'lgn rc')
lqn config = dict(layout)
lqn config['elements'] = 'lqn rc'
                                                                                                                                                                                                       nest.ConnectLayer(v1, v1,
lgn = nest.CreateLayer(lgn config)
                                                                                                                                                                                                                  'sources': {'model': 'l4 pyr'},
                                                                                                                                                                                                                  'targets': {'model': 'l4 inh'},
 # - V1 ------
                                                                                                                                                                                                                   'mask': {'circular': {'radius': 1.0}},
                                                                                                                                                                                                                   'kernel': {'linear': {'c': 1.0, 'a': -1.0}},
nest.CopyModel('iaf neuron', 'l4 pyr')
                                                                                                                                                                                                                  'delays': 1.0,
nest.CopyModel('iaf neuron', 'l4 inh')
                                                                                                                                                                                                                  'weights': 2.0})
nest.CopyModel('iaf neuron', 'l6 pyr')
                                                                                                                                                                                                       # - V1/L4 -> V1/L6 -----
v1 config = dict(layout)
v1 config['elements']
         \begin{array}{c} & = \left[ \left( 14 \text{ pyr} \right)^{2} \right)^{2} + \left( 14 \text{ inh}^{2} \right)^{2} \\ \text{Oldenburg 6 April 2009} & \left( 14 \text{ inh}^{2} \right)^{2} \right)^{1} + \left( 116 \text{ pyr}^{2} \right)^{2} \\ & \otimes \left( 14 \text{ mh}^{2} \right)^
v1 = nest.CreateLayer(v1 config)
```

```
nest.ConnectLaver(retina, lgn,
  {'connection type': 'convergent',
   'mask': { 'grid': {'rows': 1, 'columns': 1} },
# - LGN -> V1/L4 -----
nest.ConnectLaver(lqn, v1,
  {'connection type': 'convergent',
   'targets': {'model': 'l4 pyr'},
   'mask': {'rectangular':
           {'lower left': [-0.4, -0.1],
            'upper right': [0.4, 0.1]}},
   'delays': {'uniform': {'min': 2, 'max': 3}}})
# - V1/L4 -> V1/L4 ------
  {'connection type': 'divergent',
```

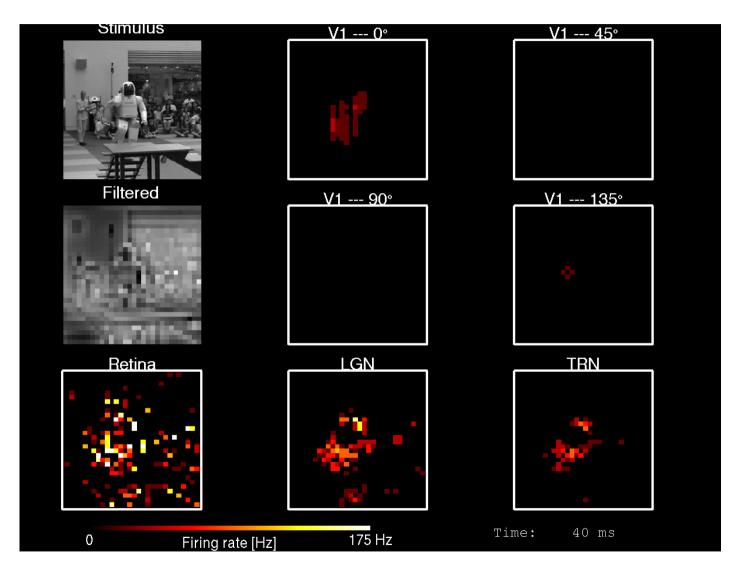
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Showtime

Simplified version of Hill & Tononi (2005)



Larger visual-pathway model



How "science" is neuronal network simulation today?

Scientific method

- Thorough critique of methods, observations, and conclusions
- Validation based on independent reproduction
- Accumulation of knowledge through exchange, evolution and (sometimes) revolution of ideas
- Requires precise and comprehensible description of research

Reproducibility

Neuronal network simulations are generally not reproducible today (and everyone knows ...)

Example 1

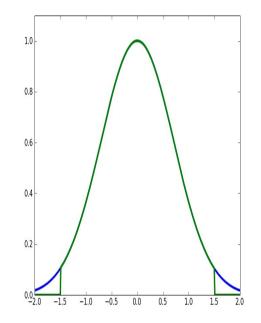
- Single neuron model
- Generally well presented
- Paper-and-pencil analysis shows that row "3" should have no spikes
- Could not be resolved in collaboration with author
- Probably figure mix-up
- No qualitative consequences

Example 2

- Well-known integrate-and-fire network model
- Chosen as benchmark for simulator comparison
- Author of paper unable to reproduce figures from his own paper with his own simulator
- Differences probably due to "minor" changes in simulator code
- Never resolved
- No qualitative consequences

Example 3

- Well-known paper on plasticity
- Neuronal connections based on Gaussian profile
- Reproduction failed qualitatively
- Inspection of original C-code revealed Gaussian with cut-off
- Were original authors aware of role of cut-off?

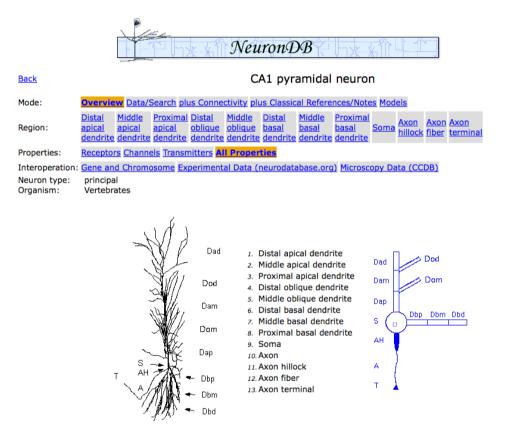


How bad is it?

- "It is currently impossible to reproduce and validate most of the results that computational scientist publish ..." (Stodden, 2009)
- "[A]n article about computational science in a scientific publication is not the scholarship itself, it is merely advertising on scholarship." (J Claerbout)
- Shooting-star crystallographer had to retract six papers because "a homemade data-analysis program had flipped two columns of data" (Science 314:1856 2006).
- See Victoria Stodden for more (http://www.stanford.edu/~vcs/)

Sharing and re-use

Resources: NeuronDB & ModelDB





CA1 pyramidal neuron (Migliore et al 1999)

Accession: 2796

Hippocampal CA1 pyramidal neuron model from the paper M.Migliore, D.A Hoffman, J.C. Magee and D. Johnston (1999) Role of an A-type K+ conductance in the back-propagation of action potentials in the dendrites of hippocampal pyramidal neurons, J. Comput. Neurosci. 7, 5-15. Instructions are provided in the below README file.Contact michele.migliore@pa.ibf.cnr.it if you have any questions about the implementation of the model.

Reference: Migliore M, Hoffman DA, Magee JC, Johnston D (1999) Role of an A-type K+ conductance in the back-propagation of action potentials in the dendrites of hippocampal pyramidal neurons. <u>J Comput</u> Neurosci 7:5-15 [PubMed]

Citations Citation Browser

Model Information (C	lick on a link to find other models with that property)
Model Type:	Neuron or other electrically excitable cell;
Brain	
Region(s)/Organism:	
Cell Type(s):	CA1 pyramidal neuron;
Channel(s):	<u>I Na,t; I A; I K;</u>
Gap Junctions:	
Receptor(s):	
Transmitter(s):	
Simulation	
Environment:	
Model Concept(s):	Dendritic Action Potentials; Active Dendrites; Detailed Neuronal Models:
Implementer(c):	Migliore, Michele ;
implementer(s):	Migliore, Michele;
Search NeuronDB for i	information about: CA1 pyramidal neuron; I A; I K; I Na,t;
Model files Downloa	d zip file Auto-launch Help downloading and running mod
	Hippogrammal CM1 purramidal nouron model from the paper

Hoder mes	nep downloading and running models
07	Hippocampal CA1 pyramidal neuron model from the paper M.Migliore, D.A Hoffman, J.C. Magee and D. Johnston
🗅 <u>ca1</u>	Role of an A-type K+ conductance in the back-propagation of

But are they used?

• Google Scholar search for "ModelDB Accession Number":

Analog Modulation of Mossy Fiber Transmission Is Uncoupled from Changes in Presynaptic Ca2+ -
ineurosci.org
R Scott, A Ruiz, C Henneberger, DM Kullmann, DA ... - Journal of Neuroscience, 2008 - Soc Neuroscience
... dentate granule cell adapted from the study by Schmidt-Hieber et al., 2007 Go)
(cell number 7, imported from SenseLab; ModeIDB accession number 95960, http ...
Sitert av 1 - Beslektede artikler - Websøk - Alle 2 versjoner

A modeling study suggesting a possible pharmacological target to mitigate the effects of ethanol on ... M Migliore, C Cannia, CC Canavier - Journal of Neurophysiology, 2008 - Am Physiological Soc Beslektede artikler - Websøk - Alle 4 versjoner

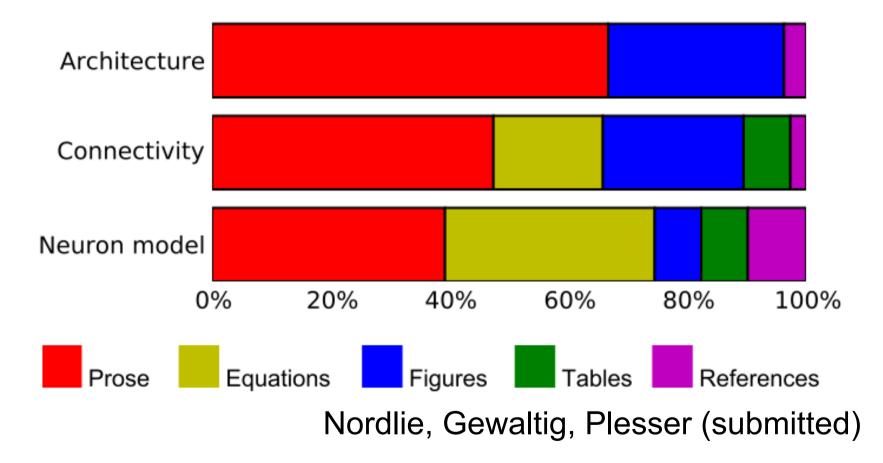
Translating network models to parallel hardware in NEURON - > psu.edu [PDF] - BIBSYS eText ML Hines, NT Carnevale - Journal of Neuroscience Methods, 2008 - Elsevier Sitert av 2 - Beslektede artikler - Websøk - Alle 3 versjoner

Improved Focalization of Electrical Microstimulation Using Microelectrode Arrays: A Modeling Study - BIBSYS eText S Joucla, B Yvert - PLoS ONE, 2009 - pubmedcentral.nih.gov ... both a straight fiber (Figure 9.A) and a complex CNS neuron, taken from the literature [32] and obtained from ModelDB (accession number 2448) (Figure 9.B). The ... Websøk

Why not?

- Little tradition for use of standard tools/simulators
- Difficult to port models from one simulator to other
- PhD students like to write their own simulators
 - Instant gratification from software development
 - Unaware of pitfalls
 - Desire for "total understanding & control"
 - Lack of compentence among supervisors?
- Models described in widely different ways in literature
- Rarely ever in a way facilitating re-building

Ways of model description



Description vs Development

- Models often described with detailed references to biological literature
- Biological justification may obfuscate concise of resulting model architecture
- But in fact
 - Design decisions often based on "what was needed to make model work"
 - Some decisions motivated by external factors, eg need to define student project
- Last to points rarely mentioned in papers
- Information important for re-use lacking Oldenburg 6 April 2009 © H E Plesser / UMB

Perspectives

- Increasing awareness of advantages of "standard simulators" (Neuron, NEST, Genesis/ Moose, PCSim, Brian)
- Review by Brette et al, J Comp Neurosci 2007
- Simulator integration (PyNN, Music)
- SBML & CellML showing advantages of standards for sharing models
- INCF Task Force on Standard Language for Neuronal Network Models
- BUT: Requires conscious effort by all in the field