

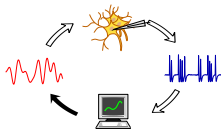
***RELACS* —** **a modular software platform for closed-loop experiments**

Jan Benda

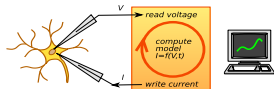
Biozentrum Martinsried
Ludwig-Maximilians Universität München



Content



Closed-loop experiments



Dynamic clamp

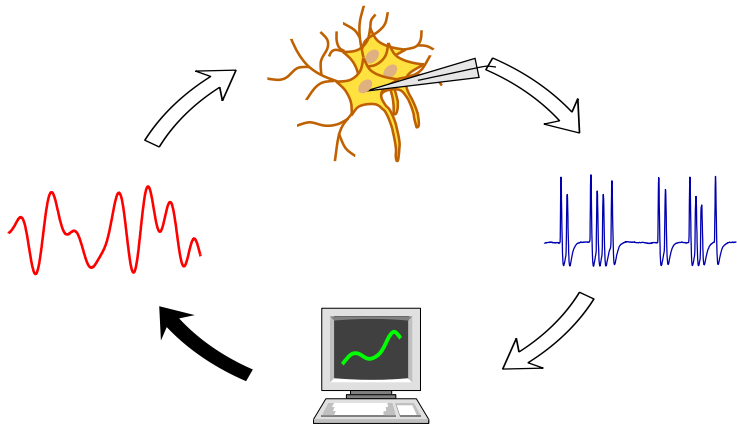


RELACS



Metadata

Closed-loop experiments



“Traditional” experiments

1. A set of stimuli and a more or less fixed experimental protocol are prepared

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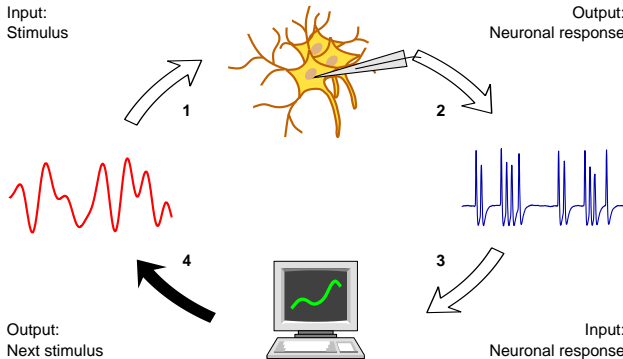
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3. The data are analyzed offline
4. The stimuli and the protocol are modified
5. A new set of recordings is made
6. After several iterations a paper is written

Closed-loop experiments

1. Present a stimulus
2. Record the response
3. Immediately analyze and visualize the data
4. Generate the next stimulus

Input:
Stimulus

Output:
Neuronal response



Simple closed-loop experiments

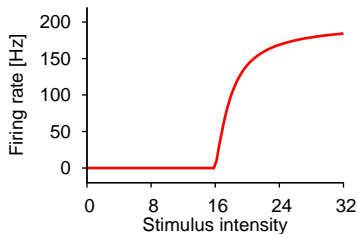
- Online visualization of processed data:
 - General infos, e.g. quality of spike detection, sensitivity of the cell, temperature, condition of animal, ...
 - Specific results, e.g. spike raster, firing rates, spike-triggered averages, ...
- ⇒ Speeds up manual (“traditional”) closed-loop

Simple closed-loop experiments

- Online visualization of processed data:
 - General infos, e.g. quality of spike detection, sensitivity of the cell, temperature, condition of animal, ...
 - Specific results, e.g. spike raster, firing rates, spike-triggered averages, ...
- ⇒ Speeds up manual (“traditional”) closed-loop
- Set stimuli relative to the neuron’s dynamic range
- Automatically control motorized electrodes (great for dual unit recordings!)
- Optimize tuning curve measurements
- ...

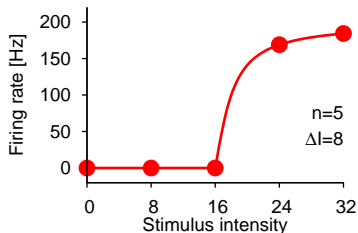
Example: tuning curve measurement

Traditional:



Example: tuning curve measurement

Traditional:

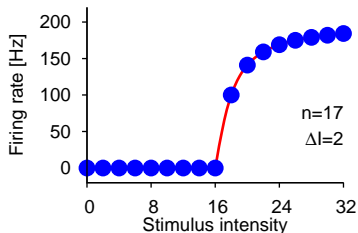


either:

fast \rightarrow low resolution

Example: tuning curve measurement

Traditional:



either:

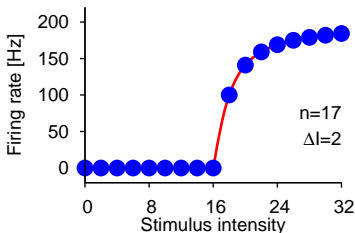
fast \rightarrow low resolution

or:

high resolution \rightarrow slow

Example: tuning curve measurement

Traditional:



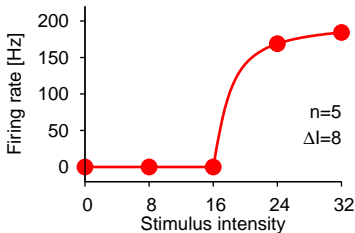
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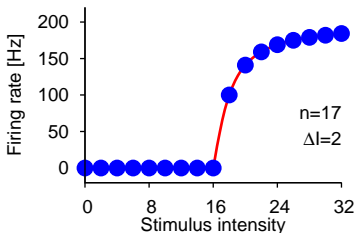
Closed loop:



1. start with low resolution

Example: tuning curve measurement

Traditional:



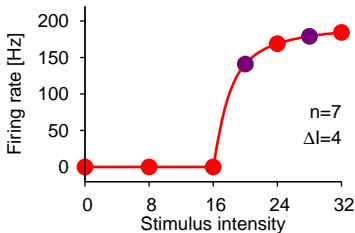
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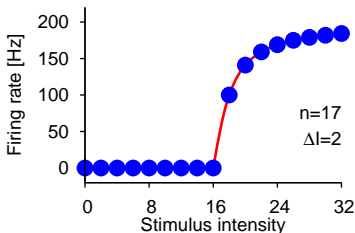
Closed loop:



1. start with low resolution
2. increase resolution where necessary!

Example: tuning curve measurement

Traditional:



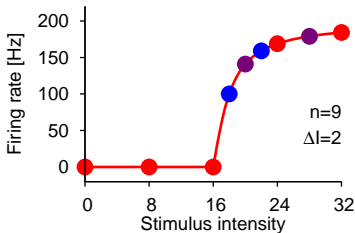
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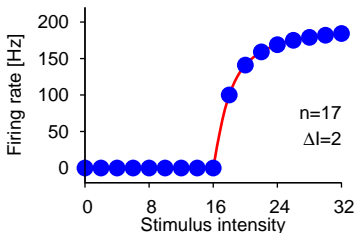
Closed loop:



1. start with low resolution
2. increase resolution where necessary!
3. further increase resolution

Example: tuning curve measurement

Traditional:



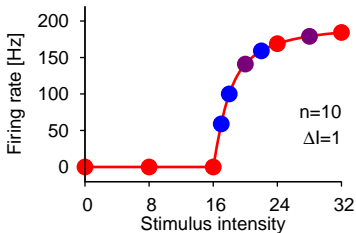
either:

fast \rightarrow low resolution

or:

high resolution \rightarrow slow

Closed loop:



1. start with low resolution
2. increase resolution *where necessary!*
3. further increase resolution

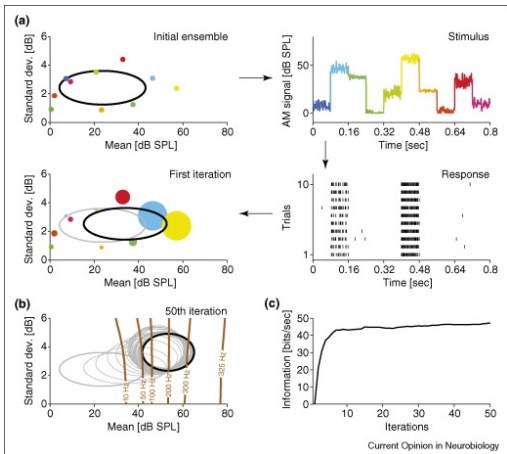
Advanced closed-loop experiments

New experimental designs are possible:

- Optimal search for a neuron's receptive field.
- Search for stimuli that drive a neuron in an "optimal" way.
- Find set's of stimulus parameter that result in the same response (iso-response method).
- ...

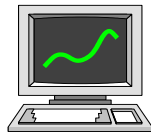
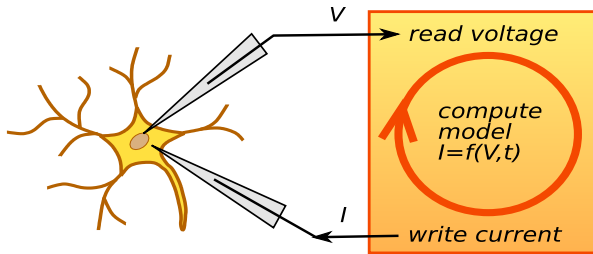
Benda et al. (2007): "From response to stimulus: adaptive sampling in sensory physiology." *Curr. Opin. Neurobiol.* **17**: 430–436.

Example: optimal stimulus ensembles



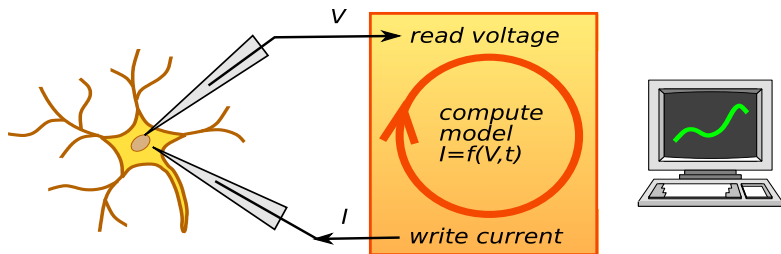
Machens et al. (2005) *Neuron* 17: 47–56.

Dynamic clamp



Dynamic clamp

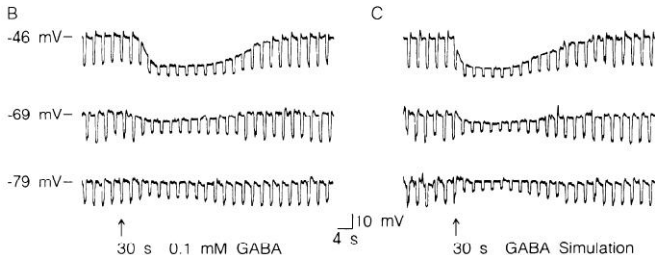
Current-clamp, with the current I computed as a function of the measured membrane potential V .



Closed-loop at a per sample time scale (tens of kHz).

Artificial conductances

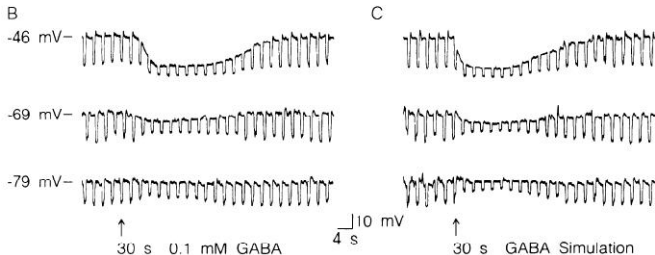
$$I = g(t) \cdot (V - E)$$



Andrew A. Sharp, Michael B. O'Neil, L. F. Abbott, & Eve Marder (1993) *J Neurophysiol*

Artificial conductances

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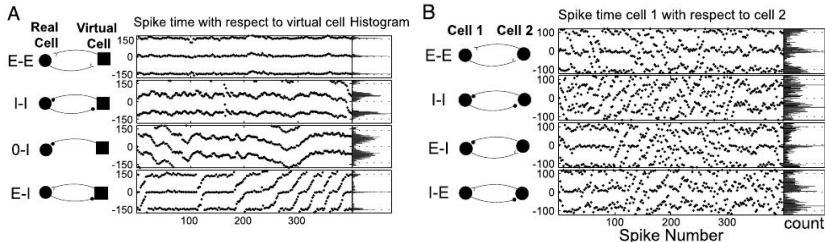
Andrew A. Sharp, Michael B. O'Neil, L. F. Abbott, & Eve Marder (1993) *J Neurophysiol*

- Synaptic conductances
- Voltage-gated conductances

Artificial networks

$$I_1 = g_{syn}(V_2) \cdot (V_1 - E)$$

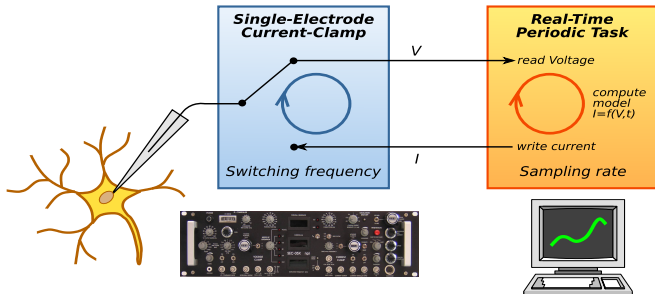
$$I_2 = g_{syn}(V_1) \cdot (V_2 - E)$$



Theoden I. Netoff, Matthew I. Banks, Alan D. Dorval, Corey D. Acker, Julie S. Haas, Nancy Kopell, & John A. White (2005) *J Neurophysiol*

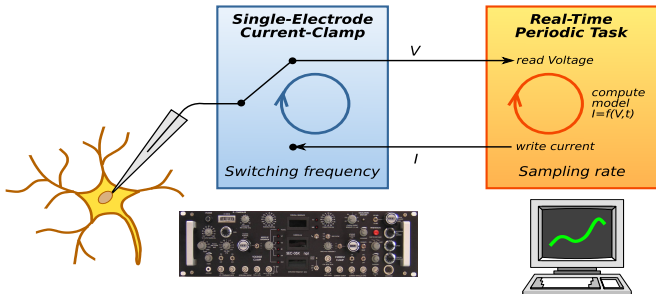
- Artificially couple real neurons
- Couple with simulated neurons

Discontinuous CC and dynamic clamp



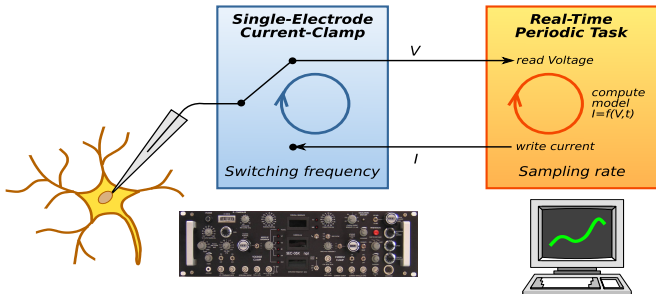
- Sampling rate \leq switching frequency/2

Discontinuous CC and dynamic clamp



- Sampling rate \leq switching frequency / 2
- ⇒ Synchronize switching and dynamic clamp cycles!
- Sampling rate = switching frequency
- (in collaboration with R. Polder, npf electronic)

Discontinuous CC and dynamic clamp



- Sampling rate \leq switching frequency / 2
- \Rightarrow Synchronize switching and dynamic clamp cycles!
Sampling rate = switching frequency
(in collaboration with R. Polder, npf electronic)
- NPI SEC: variable switching frequency $\gg 10$ kHz
independent of C compensation

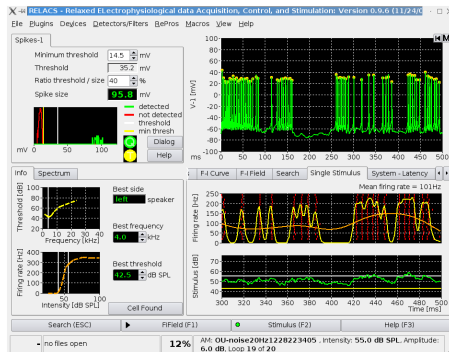
relacs_____

... enjoy your recordings



www.relacs.net

Relaxed Electrophysiological data Acquisition, Control, and Stimulation
RELACS is a framework for closed-loop experiments

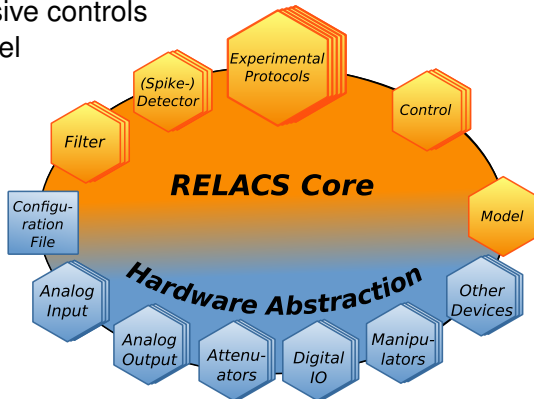


⇒ currently 13 scientific publications based on
RELACS data in *Neuron*, *J Neurosci*, *PLoS Biol*,
Nat Neurosci, *J Neurophysiol*, etc.

Modular design

RELACS core with flexible C++ Plugins for

- hardware abstraction
- data pre-processing (filter, spike detectors)
- **experimental protocols**
- passive controls
- model

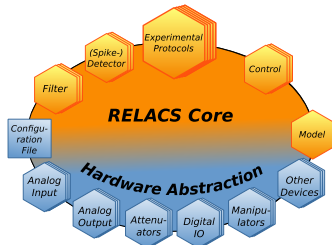


Hardware independent protocols

RELACS integrates all hardware components.

Experimental protocols for *RELACS*

- are implemented independently of specific hardware
- can be used on all the different experimental setups in your lab without any modifications
- can be **shared** with other labs



Free and open source software

RELACS is open source and free software distributed under the GNU General Public License (GPL).

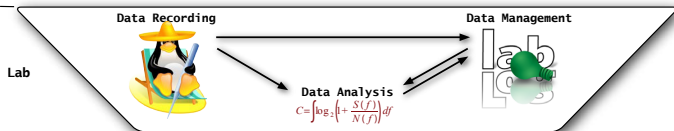
- No hassle with licenses of commercial software.
- Add whatever new feature you need directly to the program.
- Share the program and your specific experimental protocols with your collaborators.
- Know what the data-analysis algorithms are doing!

Talking about data

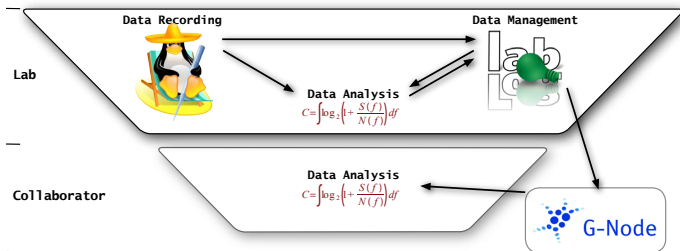
an extensible framework
for metadata exchange



The data-chain



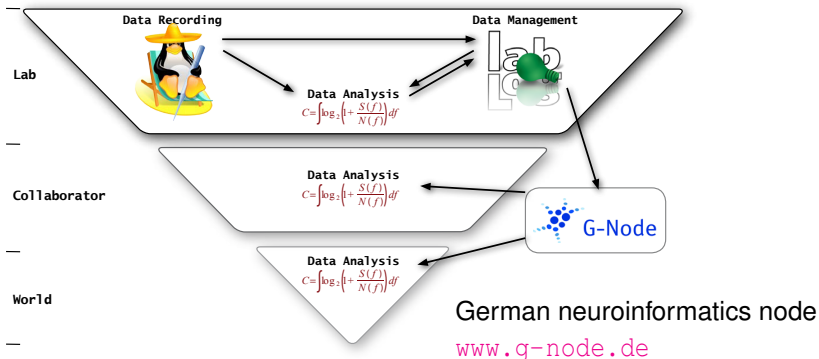
The data-chain



German neuroinformatics node

www.g-node.de

The data-chain



- All data transfer requires talking about data.
- How to exchange metadata?

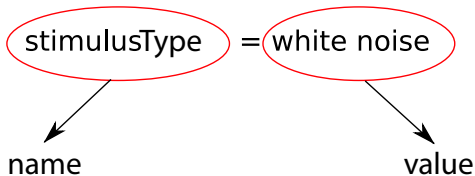
Metadata

- is “data about data”.
- describe recording conditions.
- essential for data analysis, management, and sharing.

stimulusType = white noise

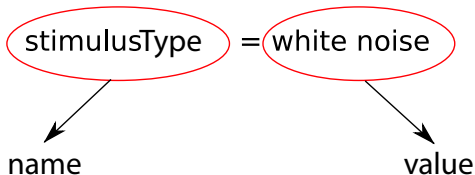
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The metadata problem

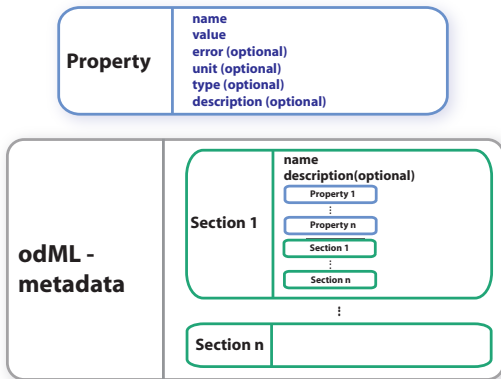
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- What name to choose?
- What does it mean?
- How to organize metadata?

odML — open metadata markup language

Structure:



Implemented as the [odML XML Schema](#)

odML — open metadata markup language

Vocabularies: names & definitions

HardwareSettings:

Amplifier:

name	type	description
Gain	float	The amplifier gain.
HighpassCutoff	float	The cutoff frequency of the amplifier's highpass filter. Given in Hz.
LowpassCutoff	float	The cutoff frequency of the amplifier's lowpass filter. Given in Hz.
Mode	string	The amplifier mode. E.g. Bridge, CC, VC etc.

How to use odML?

1. Assemble properties:

- If you find an appropriate property in the [odML](#)-vocabularies, use it!
- Ignore all properties that do not match.
- Add your own properties that are not yet in the vocabulary, if possible with a description.

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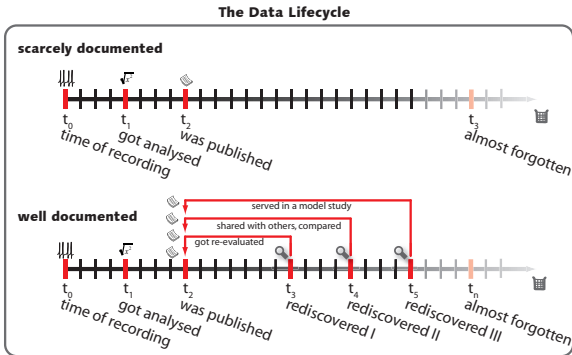
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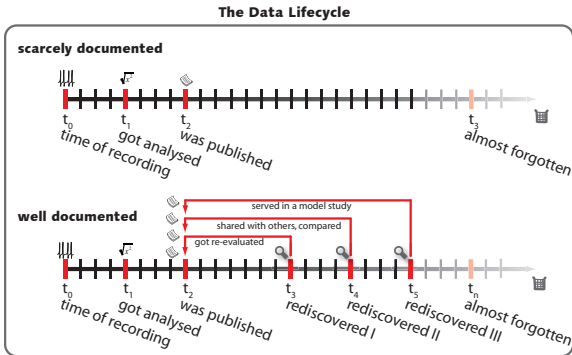
3. Transfer them to an analysis or database program

The data life-cycle



- Meta information tends to vanish with time.
- Thus, re-using of old data is a tedious business.
- Data should be annotated as early as possible (preferentially at the time of acquisition, e.g. with *RELACS*).

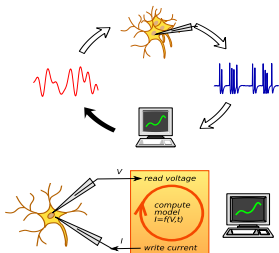
The data life-cycle



- odML provides a simple and flexible standard
- Well annotated data can be found and reused easily

⇒ Your data deserves it!

Summary



Closed-loop experiments

Novel experimental designs

Dynamic clamp

Artificial conductances and hybrid networks

— R. Polder, npj electronic, Tamm

RELACS www.relacs.net

Software platform for closed-loop and dynamic clamp experiments

— NeurOnline: S. Rotter, M. Ambard, A. Brandt, C. Boucsein, Freiburg

Metadata — odML

A standard for sharing data

— J. Grewe & G-node, LMU Munich

Experimental protocol example

```
int Example::main( void ) {
    // some initialization ...
    OutData signal;
    signal.setTrace( "LeftSpeaker" );
    signal.sineWave( frequency, duration, amplitude );
    SampleDataD rate( 0.0, duration, 0.001 );
    for ( int counter=0; counter<Repeats; counter++ ) {
        write( signal );
        sleep( duration + pause );
        EventData spikes( events( SpikeEvents[0] ),
                           events( SpikeEvents[0] ).signalTime(),
                           events( SpikeEvents[0] ).signalTime() + duration );
        double meanrate = spikes.rate( 0.3*duration, duration );
        spikes.addRate( rate, counter, GaussKernel( sigma ) );
        P.clear();
        P.plot( rate, 1000.0, Plot::Yellow, 2, Plot::Solid );
        P.draw();
        if ( meanrate < targetrate ) {
            amplitude *= 2.0;
            signal.sineWave( frequency, duration, amplitude );
        }
    }
}
```

RELACS C++ library for data analysis

Data structures (classes, container):

- *Array* — Basic 1-D vector
- *SampleData* — 1-D data vector with regularly sampled time axis
- *Map* — Sequence of $x|y$ data pairs

Algorithms:

- basic statistics (moments, quartiles, histogram)
- power spectra, coherence, transfer function
- linear fits
- non-linear fits (Simplex, Levenberg-Marquardt)

RELACS C++ library for data analysis

Data structures (classes, container):

- *EventData* — Spikes and other point process data
- *EventList* — Multi-trial spike trains

Algorithms:

- firing rates (mean, PSTH binned/kernel, 1/ISI)
- CV, Fano factor, ISI correlation
- vector strength, reliability, jitter
- mutual information (lower and upper bound)

Current odML developments

Done:

- Schema definition converges to version 1.
- Java and MatLab library to read, write and manipulate odML-files.
- Editor for odML metadata.

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- Schema definition converges to version 1.
- Java and MatLab library to read, write and manipulate odML-files.
- Editor for odML metadata.

On the way:

- Definition of the vocabularies.

Planned:

- Libraries for C/C++, Python ...
- collaboration with various initiatives (CRCNS, NIF, etc.)

odML — Details

odML-Property:

element name	description	occurrence
name	The name of this property.	1
value	The value of this property.	1 - ∞
error	An error estimate of the value(s).	0 - ∞
unit	The unit of the value(s).	0 - 1
type	The datatype of the value e.g. int, float, string, date, time, binary, etc.	1
id	An identifier for each value, e.g. for a database.	0 - ∞
nameDefinition	Defines the property.	0 - 1
valueDefinition	Defines each individual value.	0 - ∞
parent	This property is only meaningful if the parent property exists.	0 - 1
parentValue	This property is only meaningful for a specific parent value.	0 - 1

odML — Details

odML-Section:

element name	description	occurrence
name	The name of this section.	1
alias	An alias name for this section.	0 - 1
id	An identifier e.g. from a database by which the information of this section can be found.	0 - 1
definition	Defines the section.	0 - 1
vocabulary	The URI of the vocabulary which defines this section.	0 - 1
parent	This section might be meaningful only if it is child of a parent section.	0 - 1
parentURI	The URI of the parent section's definition.	0 - 1
odML-Property	A section can contain properties ...	0 - ∞
odML-Section	... and subsections.	0 - ∞