



Biological Systems Modeling & Simulation ^②

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Previous Lecture



- Biomedical Signal examples (1-d, 2-d, 3-d, ...)
- Purpose of Signal Analysis
- Noise
- Frequency domain (1-d, 2-d)
- Filtering unwanted frequencies
- Sampling
- Kalman Filter
- Recursive Algorithm

Biological Signals

- Amplitude limited
- Distorted by colored noise
- Limited length
- Non-stationary
- The underlying system is unknown

Why do we need models?

Insight

- Understanding the underlying system
- Testing bed for several hypotheses

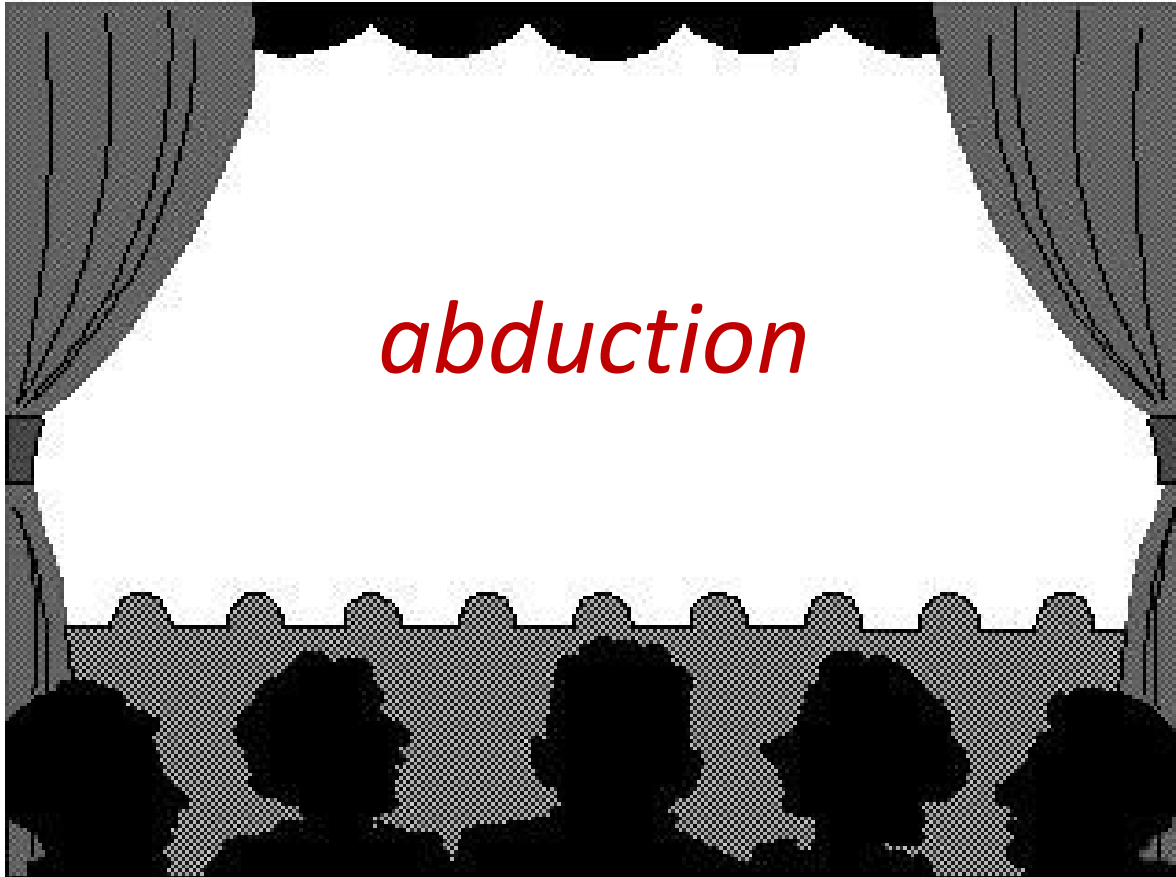
Description

- Design verification
- Explore linear behavior / non-linear nature of the biological physics (going beyond human senses)

Control

- Predictions
- Functional limits

Definition of a Model



Difficulties

1. No access to the unconditional truth
2. No unbiased observer
3. Personal opinion on a hypothesis (abduction)
4. Abduction is not an infallible way for discovering truth

Modeling a System

- **System**

(a collection of interconnected processes)

- **Model**

(a representation that approximates the behavior of an actual system)

- **Simulation**

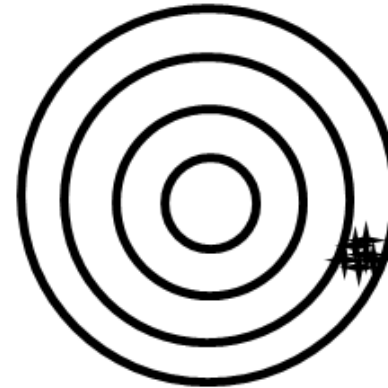
(puts the model to work)

Goodness of a Model

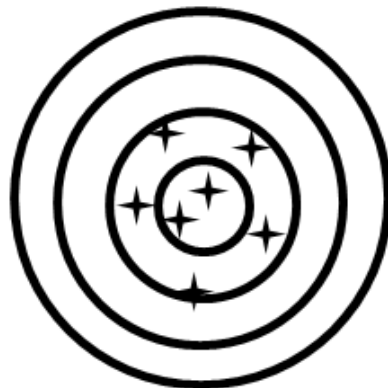
Not Accurate nor Precise



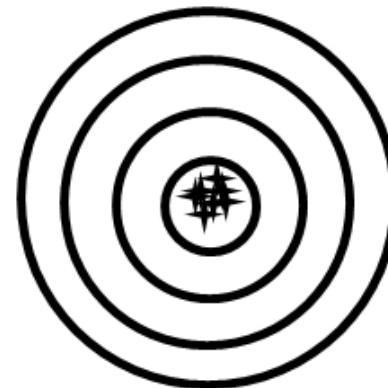
Precise but not Accurate



Accurate but not Precise



Accurate and Precise



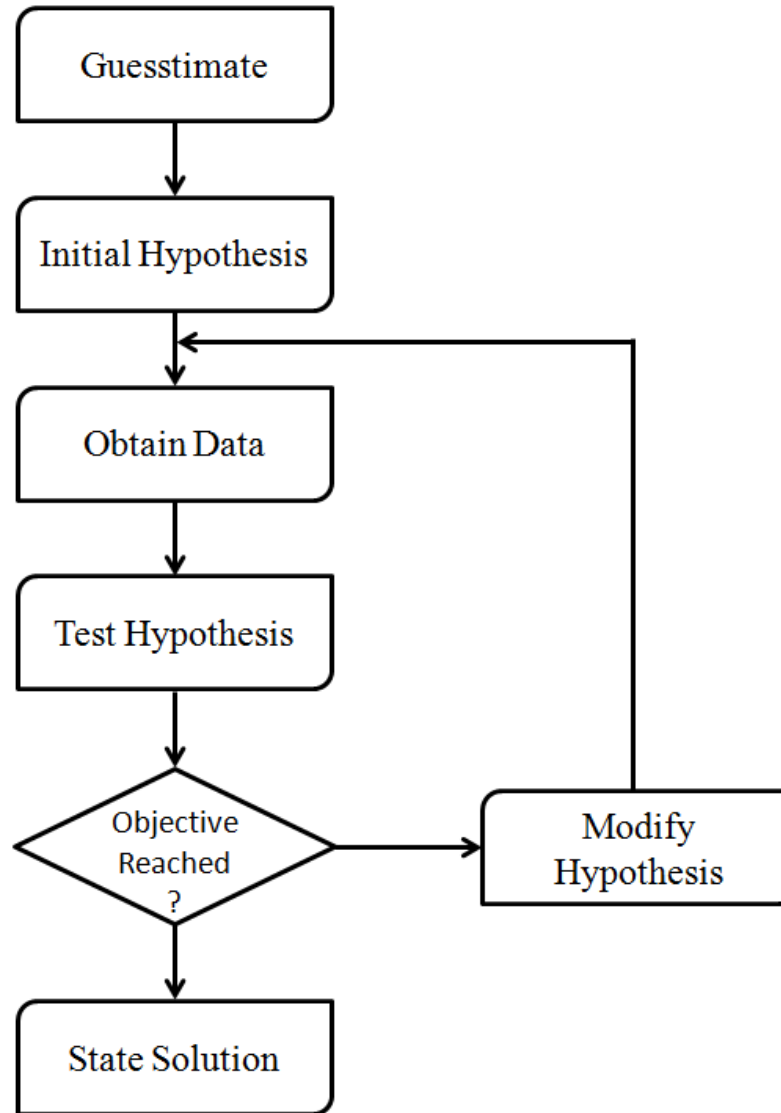
Accuracy

Difference between predicted and true value

Precision

Reproducibility of the results

Model refinement



Classification of Models

Level	Sub-cellular	Cellular	Intercellular	Tissue	Organ	Organism
Behavior		Dynamic			Static	
Chance		Deterministic			Stochastic	
Parameters		Parametric			Non-parametric	
Compartments		Single Compartment			Multiple Compartments	
Linearity		Linear			Non-Linear	
Domain		Time			Frequency	

Linear Modeling of Physiological Systems

Why linear?

- Simple to implement
- Powerful analysis tools

Transforming input into output

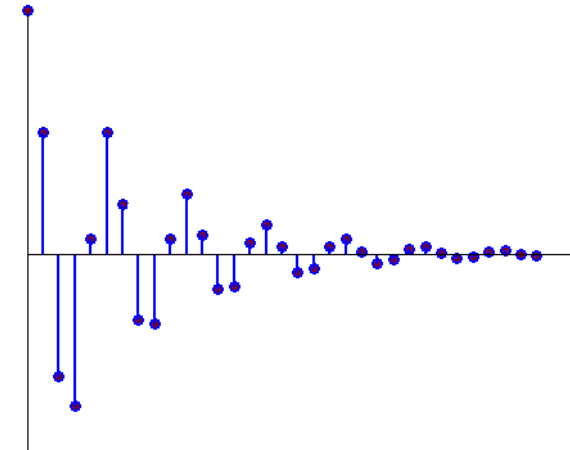
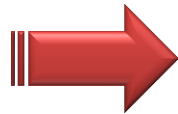
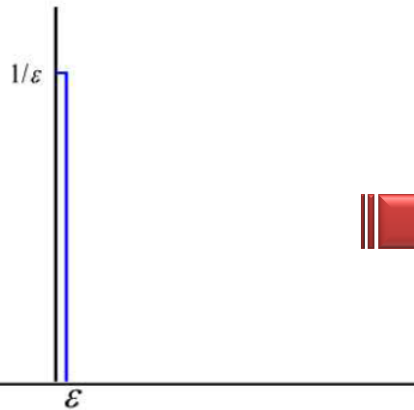
- Time variant
- Time invariant

Linear Time-invariant Systems

1. Superposition
2. Impulse Response of a system

- Testing for linearity may be done using the principle of superposition (e.g. same input at different amplitudes)
- Response of a linear system to a sinusoidal input is a sinusoid at the same frequency
- Linearity of a system means nothing without its range

Impulse Response Function

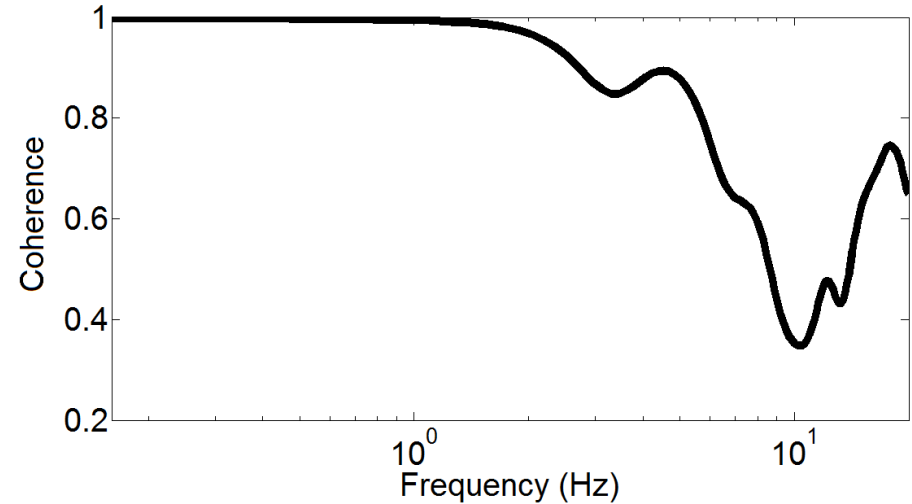
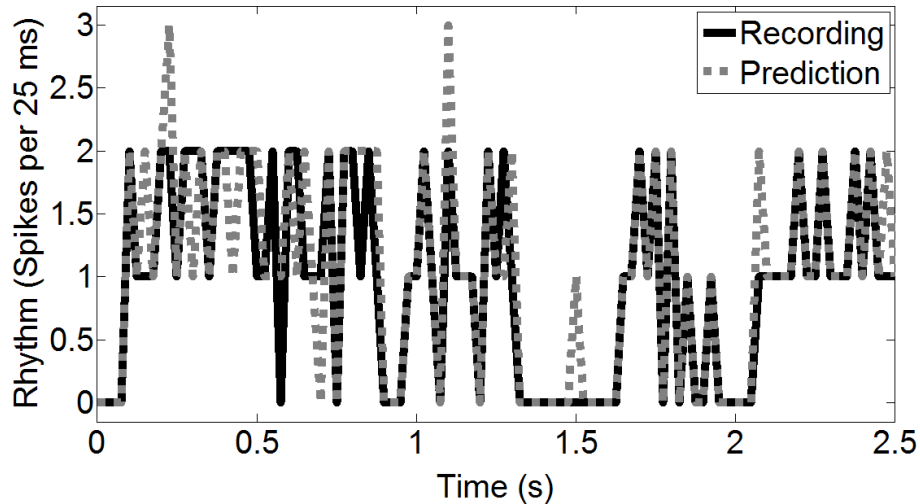


Negative Values = System anticipation

Positive Values = System memory

Living systems frequently demonstrate **predictive behavior** (e.g. visual pursuit)

Coherence



If Coherence is greater than 0 but not 1:

1. Extraneous noise is present in the measurements.
2. The system is not linear.
3. Output is due to the input as well as to other inputs.

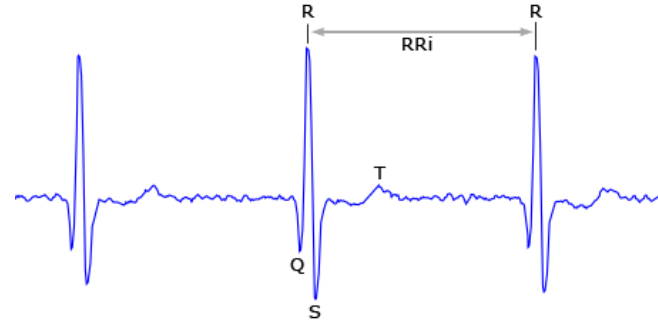
Modeling Chaos in Physiology

- Chaos ($\chi\alpha\tilde{o}\varsigma$) = preponderant of all the others
- Physiological Systems are **extremely complicated deterministic systems** for their observers
- Stochastic determinism observed in biological signals (extraordinary sensitivity to internal conditions)
- Presence of order under the absence of periodicity (**strange attractor**)
- A chaotic system is not necessarily complex

Chaos in Physiological Systems

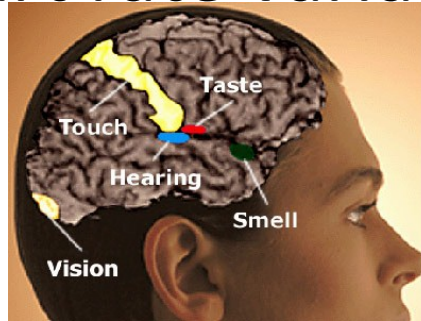
Chaos in the heart

- Sinoatrial Node
- Sympathetic & parasympathetic fibers & respiration rate → heart rate variability



Chaos in the brain

- Neuron doctrine
- Feedbacks → internal uncertainties are amplified over time



Why Chaos?

- Better **adaptation** capabilities
(e.g. heart pumps blood in various circumstances)
- **Adaptation** in the brain = learning

A never seen before stimulus in the brain, moves the brain to an un-patterned chaotic state.

- **Chaos** = normal function
Disease = acute attack of order against chaos
(e.g. the amount of chaos in the Parkinsonian brain decreases as neurons become more synchronized)