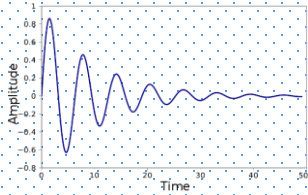
A surgeon in a white scrub cap and mask is looking at a computer monitor in an operating room. The monitor displays various medical data, including a heart rate of 62, a blood pressure of 17/65 (86), and a pulse of 17/10 (115). The text "Advanced Perioperative Monitoring. Elements of Computational Medicine." is overlaid on the image in a bold, dark green font.

# Advanced Perioperative Monitoring. Elements of Computational Medicine.

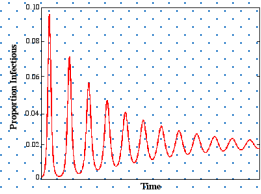
Victor Iapascurta



“In order to get an appropriate understanding of what is really happening in a body system, it is compulsory to supplement the ‘usual medical way of reasoning’ with elements of the engineering approach. As a result, along with a more knowledgeable physician (e.g. intensivist), the final beneficiary of such a combination will be the patient.

Without this it is hardly possible to speak about clinical monitoring in general and advanced monitoring in particular”

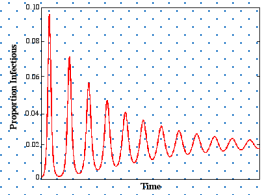
*a rule of thumb during the advanced technological era*



- **Clinician monitoring** — clinical monitoring using visual inspection, auscultation, and palpation is a primary determinant of patient safety.

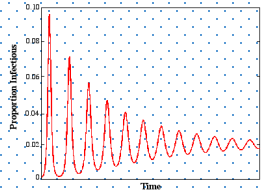
*Changes in clinical signs may be subtle, and often precede abnormalities in parameters measured by monitoring devices.*

- **Monitoring devices** do not replace clinical observation; rather, they amplify and quantify clinical information



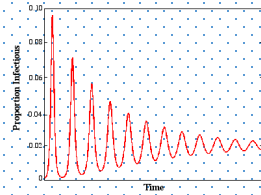
# Prerequisites & Basic Concepts

Basic knowledge concerning monitoring  
in Anesthesia and Intensive Care



## Current course

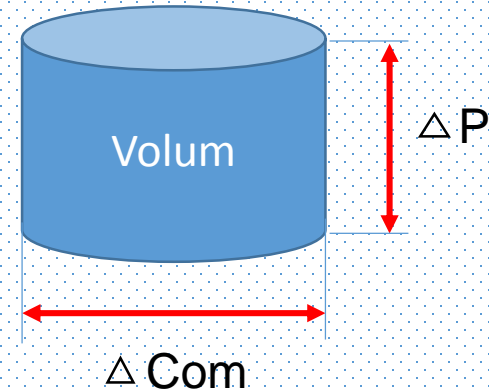
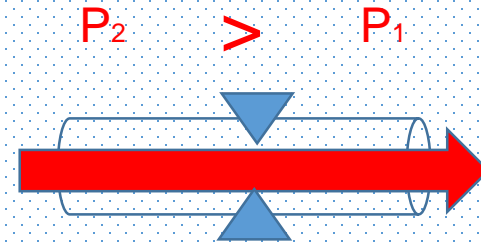
- extends the 'monitoring time frame' to the **perioperative period**
- provides additional in-depth insights based on **systemic approach** as a concept based on fundamental **interdisciplinary knowledge** which includes, but is not limited to, physics, modeling, computational sciences, etc.
- is to help the future clinician to navigate in the fast-progressing modern medical field especially concerning the emergence of **Medical Artificial Intelligence** technologies



BAASIC CONCEPTS:

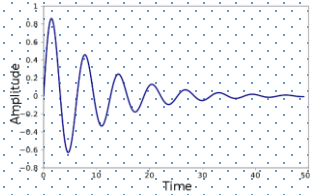
$$\text{Flow} = \frac{P_2 - P_1}{\text{Resistance}}$$

$$\Delta P = \frac{\text{Volume}}{\text{Compliance}}$$



Ohm law:  $I = \frac{U}{R}$

$$\text{Volume} = \Delta P \times \text{Compliance}; = \text{Flow} \times \text{time}$$



BAASIC CONCEPTS:

## Newton's Law of Cooling

$$\frac{dT}{dt} = k(T - T_s)$$



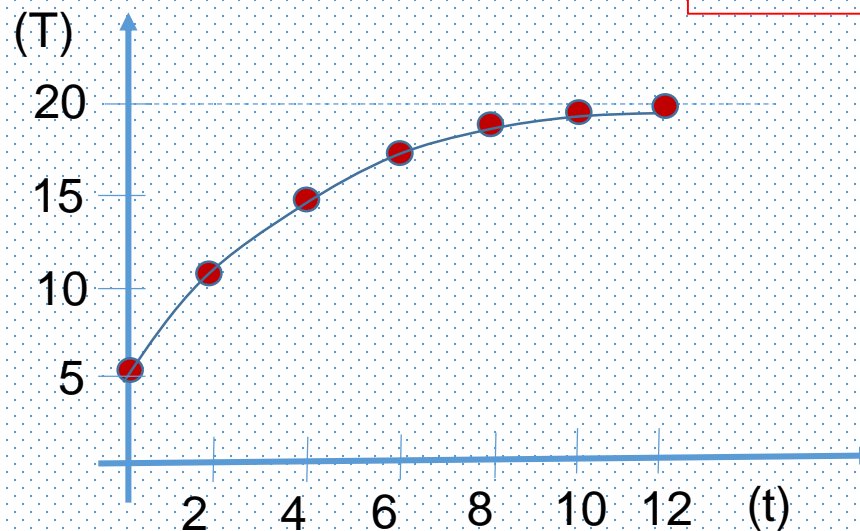
$$\frac{dT}{dt} = 0,2 (20 - T) = 0,2 \times 15 = 3 \text{ C / min}$$

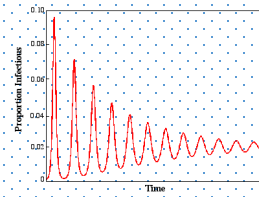
$$5 + 2 \times 3 = 11$$

$$11 + 2 \times 1,8 = 14,6$$

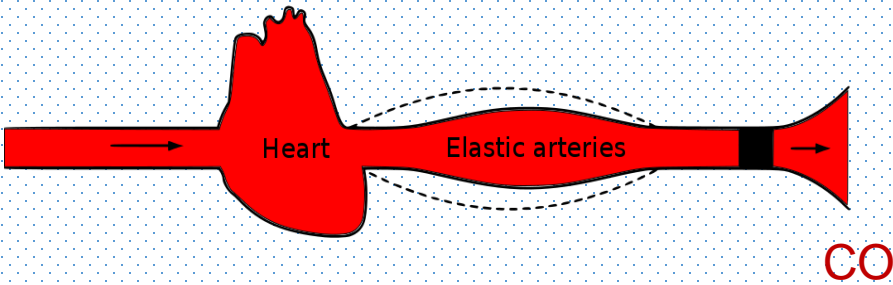
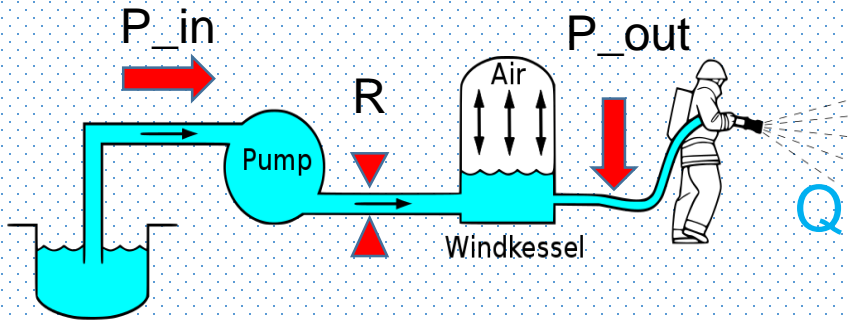
.....

- $T_0 = 5 \text{ C}$
- $T_2 = 11 \text{ C}$
- $T_4 = 14,6 \text{ C}$
- $T_6 = 16,76 \text{ C}$
- $T_8 = 18,06 \text{ C}$
- $T_{10} = 18,45 \text{ C}$
- $T_{12} = 18,76 \text{ C}$





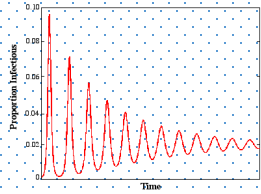
APPLYING BAASIC CONCEPTS: Windkessel Model and BP



- $Q = (P_{in} - P_{out}) / R$
- $CO = (MAP - RAP) / SVR$
- $MAP = (CO * SVR) + RAP$ 
  - $MAP = CO * SVR$
  - $CO = MAP / SVR$

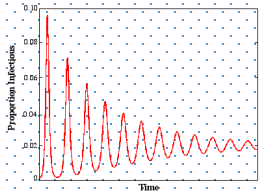
$$DO_2 = CO * 1.34 * Hb * SaO_2$$





## Clinical significance of BP (RAP/CO/SVR)

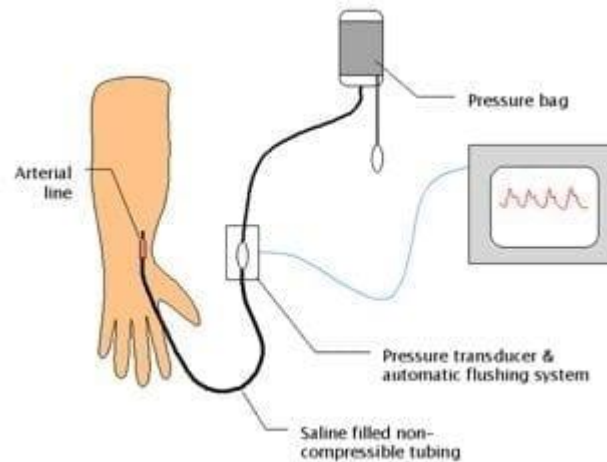
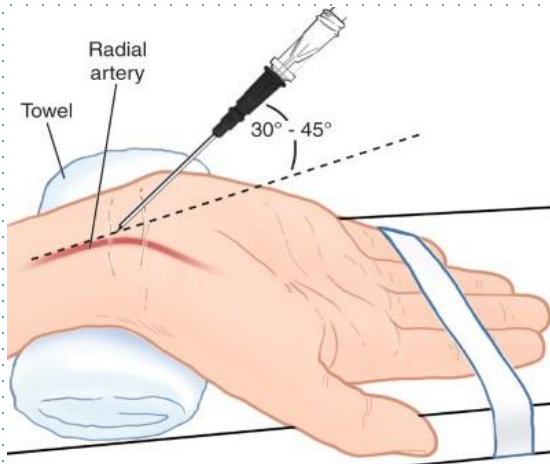
- a. **Low RAP** = hypovolemic shock.
- b. **Low CO** = cardiogenic shock
- c. **Low SVR** = vasogenic shock  
(e.g., septic shock)

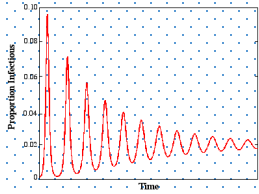


## Non invasive BP - oscillometric method

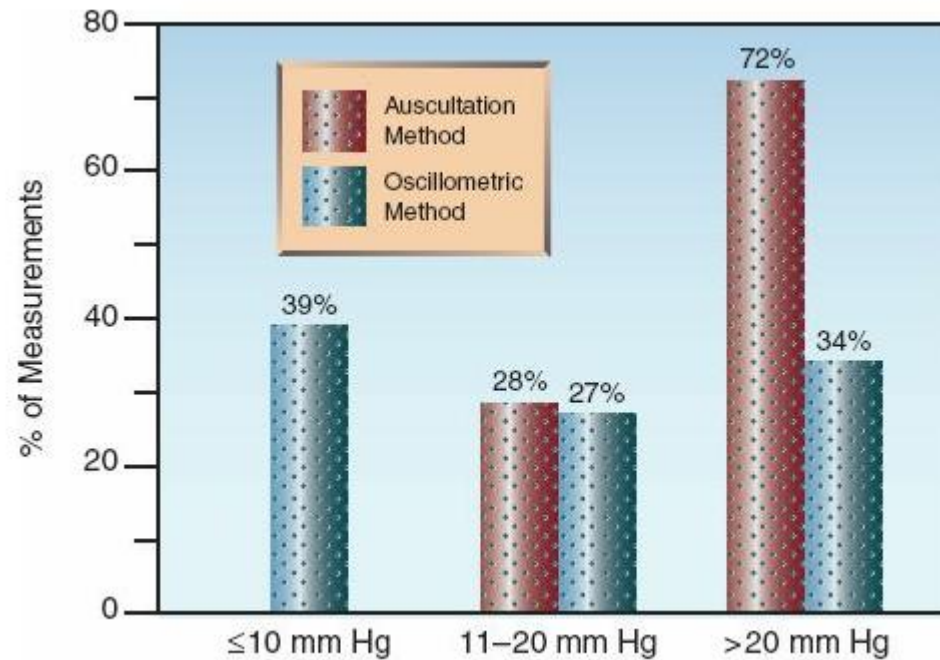


## Invasive BP - intraarterial

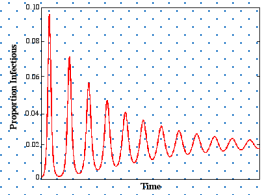




# Non invasive BP – oscillometric method highly error-prone\*



\* P.Marino, The ICU Book, 2014 e- ed, p.134 based on Bur A, Hirschl M, Herkner H, et al. Accuracy of oscillometric blood pressure measurement according to the relation between cuff size and upper-arm circumference in critically ill patients. Crit Care Med 2000; 28:371–376



## What finally counts ?

$$DO_2 = CO * 1.34 * Hb * SaO_2$$



Hemodynamic  
component



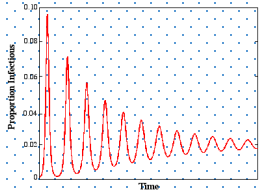
Hematic  
component



Respiratory  
component

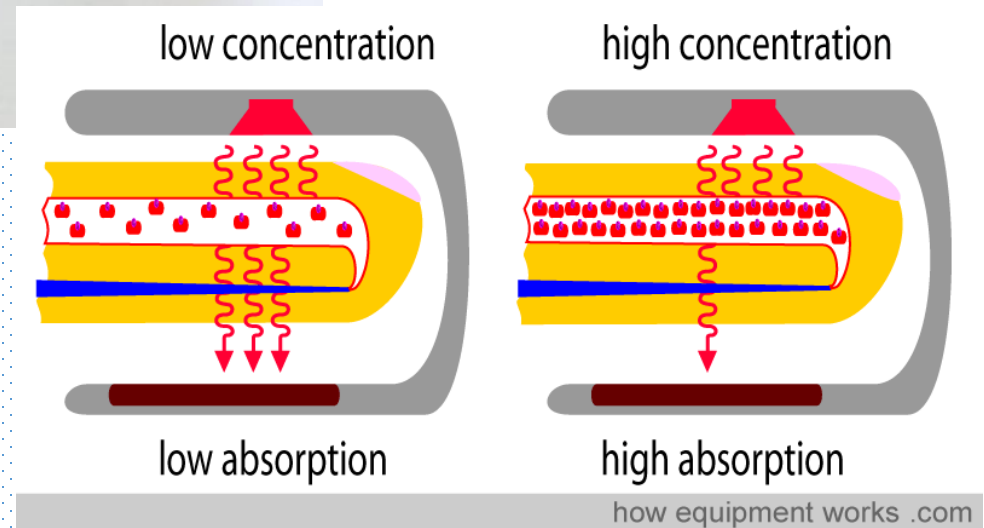
$$VO_2 = CO * 1.34 * Hb * (SaO_2 - SvO_2)$$

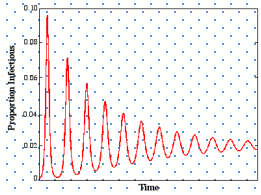
# Pulse oximetry



## SpO<sub>2</sub>

SaO<sub>2</sub> > 90 %





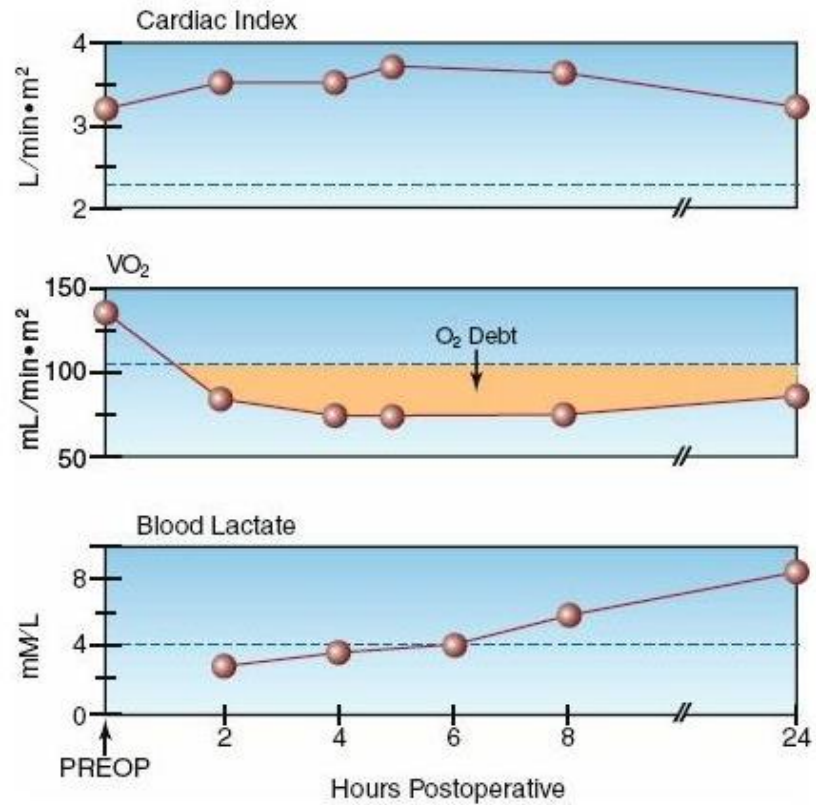
## Markers of Inadequate Tissue Oxygenation

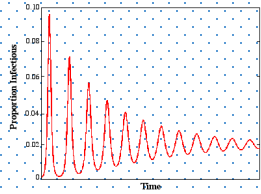
### I. Oxygen Markers

1.  $VO_2 > 200 \text{ mL/min/m}^2$  or  $< 110 \text{ mL/min/m}^2$
2.  $(SaO_2 - SvO_2) \geq 50 \%$
3.  $SvO_2 \leq 50 \%$

### II. Chemical Markers

1. Serum Lactate  $> 2 \text{ mM/L}$  (or  $\geq 4 \text{ mM/L}$ )
2. Arterial Base Deficit  $> 2 \text{ mM/L}$



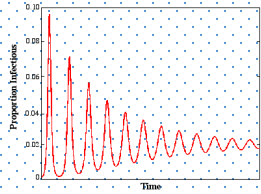


## Measuring Cardiac Output (CO)

(methods from invasive to non-invasive)

- a. Thermodilution method by Swan-Ganz catheter
- b. PiCCO, LiDCO Pulse Pressure method
- c. FloTrac method based on pulse contour analysis
- d. Thoracic Electric Bioimpedance (TEB)
- e. Doppler Ultrasound based methods

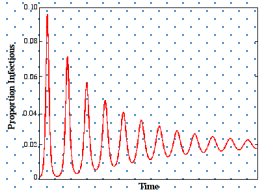
Other methods: dye dilution method, ultrasound dilution, electrical cardiometry, etc.



## Hemodynamic and Oxygen Transport Parameters

Parameter	Abbreviation	Normal Range
<b>Central Venous Pressure</b>	CVP	0 – 8 mm Hg
<b>Pulmonary Artery Wedge Pressure</b>	PAWP	5 – 15 mm Hg
<b>Cardiac Output</b>	CO	4.5 – 8 L/min
<b>Cardiac Index</b>	CI	2.7 – 4 L/min/m <sup>2</sup>
<b>Stroke Index</b>	SI	38 -60 mL/beat/m <sup>2</sup>
<b>Systemic Vascular Resistance Index</b>	SVRI	1860 – 2500 dyn*s/cm <sup>5</sup> /m <sup>2</sup>
<b>Pulmonary Vascular Resistance Index</b>	PVRI	225 – 315 dyn*s/cm <sup>5</sup> /m <sup>2</sup>
<b>O<sub>2</sub> saturation (arter.)</b>	SaO <sub>2</sub>	>90 %
<b>O<sub>2</sub> saturation (vein.)</b>	SvO <sub>2</sub>	>75%
<b>Oxygen Delivery (Index)</b>	DO <sub>2</sub>	520 – 570 mL/min/m <sup>2</sup>
<b>Oxygen Uptake (index)</b>	VO <sub>2</sub>	110 – 160 mL/min/m <sup>2</sup>
<b>Oxygen Extraction Ratio</b>	O <sub>2</sub> ER	0.2 – 0.3





# Neuromonitoring

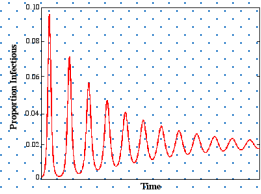
is used to assess the functional integrity of the brain, brain stem, spinal cord, or peripheral nerves.

## The goal:

- to alert the surgeon and anesthesiologist to impending injury in order to allow modification of management in time to prevent permanent damage
- to map areas of the nervous system in order to guide management
- monitoring the depth of anesthesia

## Monitoring techniques:

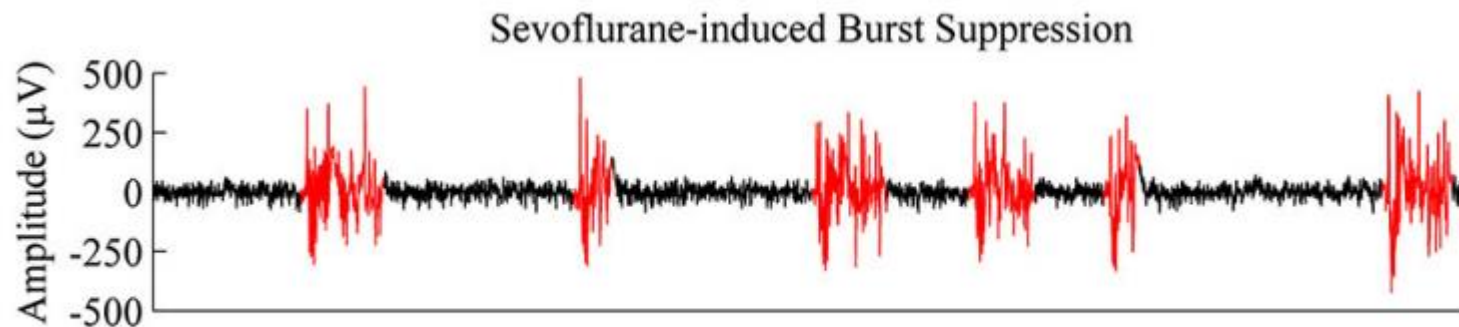
- Electroencephalography (EEG),
- Electromyography (EMG),
- Somatosensory evoked potentials (SSEPs),
- Brainstem auditory evoked potentials (BAEPs),
- Motor evoked potentials (MEPs)



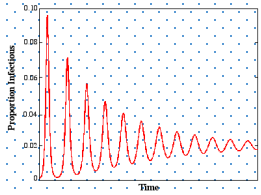
## Monitoring the Depth of General Anesthesia DGA

- Bispectral Index monitor (BIS)
- Auditory Evoked Potential monitor (AEP)
- Patient State Analyser (PSA)
- Cerebral state monitor (CSM)
- Index of Consciousness monitor (IoC)
- Entropy monitor

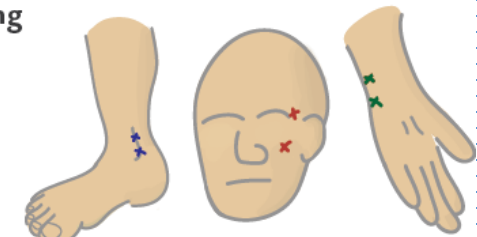
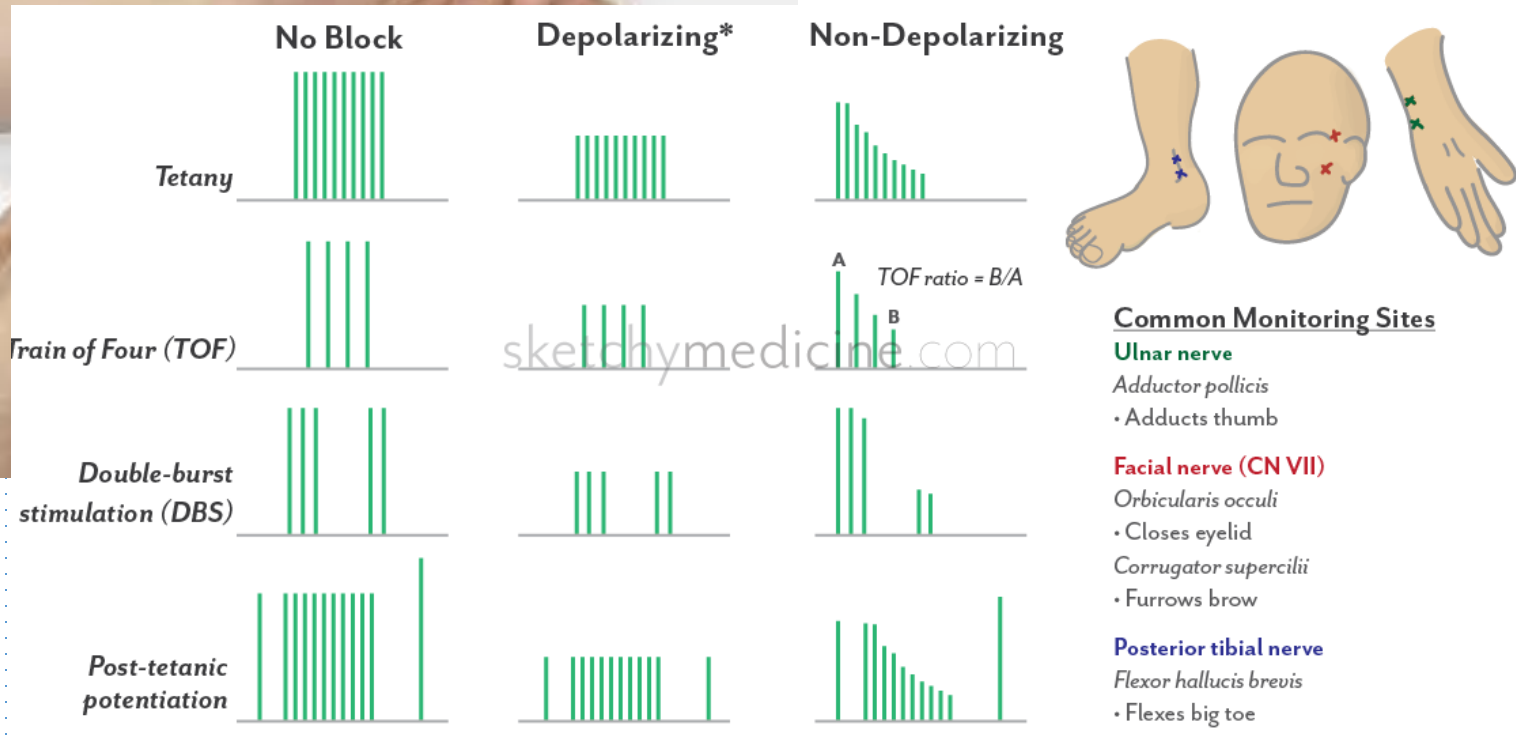
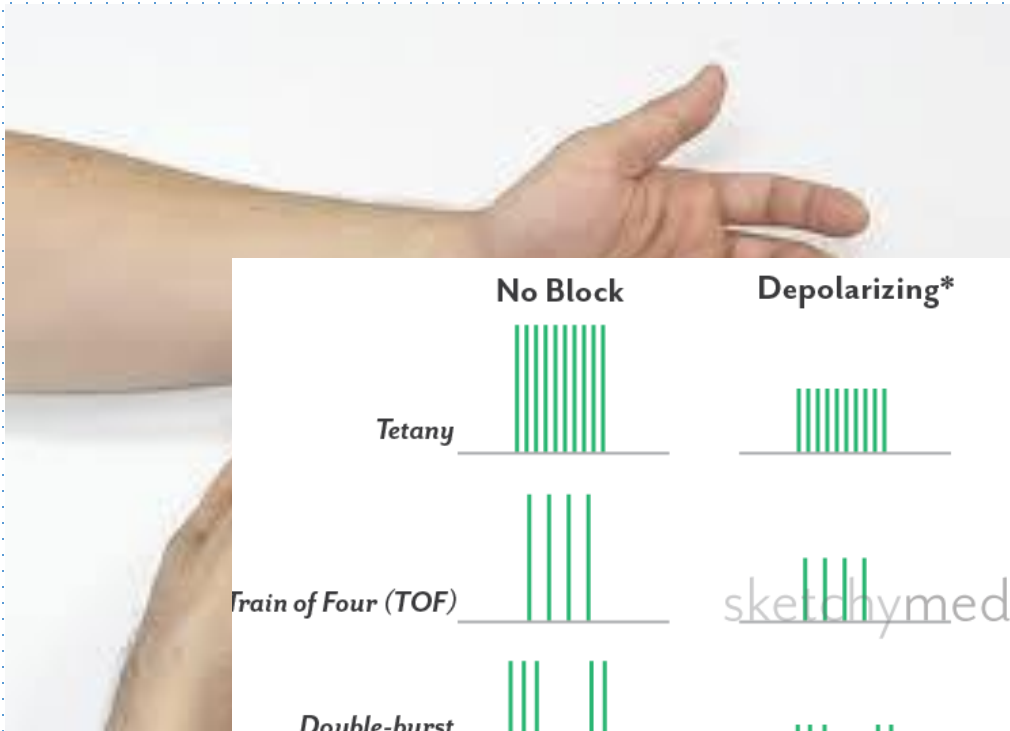
## Burst Suppression Mechanism



Jonathan D. Kenny et al., Propofol and sevoflurane induce distinct burst suppression patterns in rats, *Frontiers in Systems Neuroscience*, December 2014 | Volume 8 | Article 237



# Neuromuscular function monitoring

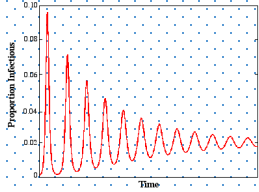


### Common Monitoring Sites

**Ulnar nerve**  
*Adductor pollicis*  
 • Adducts thumb

**Facial nerve (CN VII)**  
*Orbicularis oculi*  
 • Closes eyelid  
*Corrugator supercilii*  
 • Furrows brow

**Posterior tibial nerve**  
*Flexor hallucis brevis*  
 • Flexes big toe



# Monitoring Modalities Summary

## Respiratory System Monitoring

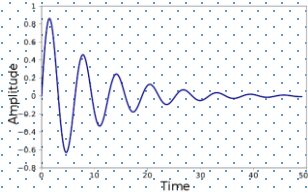
- Oxygenation (cyanosis, pulse oximetry, inspired O2 analyzer)
- Ventilation (capnography, pulmonary mechanics measurements, disconnection alarms)

## Circulatory System Monitoring

- Blood Pressure (non-invasive, invasive)
- Electrocardiogram (ECG leads, Ischemia detection, artifacts)
- Other monitoring of circulation (central venous pressure, PA catheter, CO measurement)

## Other monitoring techniques and devices

- Neuromonitoring (BIS/ AEP/PSA : DGA)
- Neuromuscular function monitoring
- Tissue oxygenation chemical markers
- Temperature monitoring
- Urine output, etc.



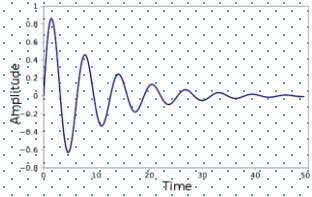
## Computational Medicine (CM)

is an emerging discipline devoted to the development of quantitative approaches for understanding the mechanisms, diagnosis and treatment of human disease through applications of mathematics, engineering and computational science.

**The core approach of CM** is to develop computational models of the molecular biology, physiology, and anatomy of disease, and apply these models to improve patient care.

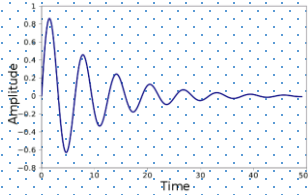
*Johns Hopkins School of Medicine*

<https://www.bme.jhu.edu/graduate/mse/degree-requirements/computational-medicine/>



## MODEL THINKING IN MEDICINE / A&IC





## MODEL THINKING IN MEDICINE / A&IC

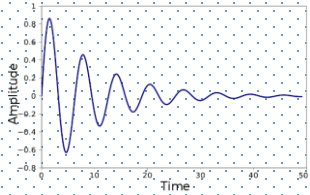
**Model** = simplified representation or abstraction of reality, that highlights important aspects, at the price of ignoring other aspects.

*“Truth ... is much too complicated to allow anything but approximations“*

***John von Neumann***

*“All models are wrong, but some are useful“*

***J.P. Box***

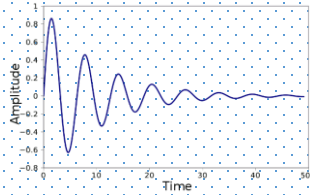


## MODEL THINKING IN MEDICINE / A&IC

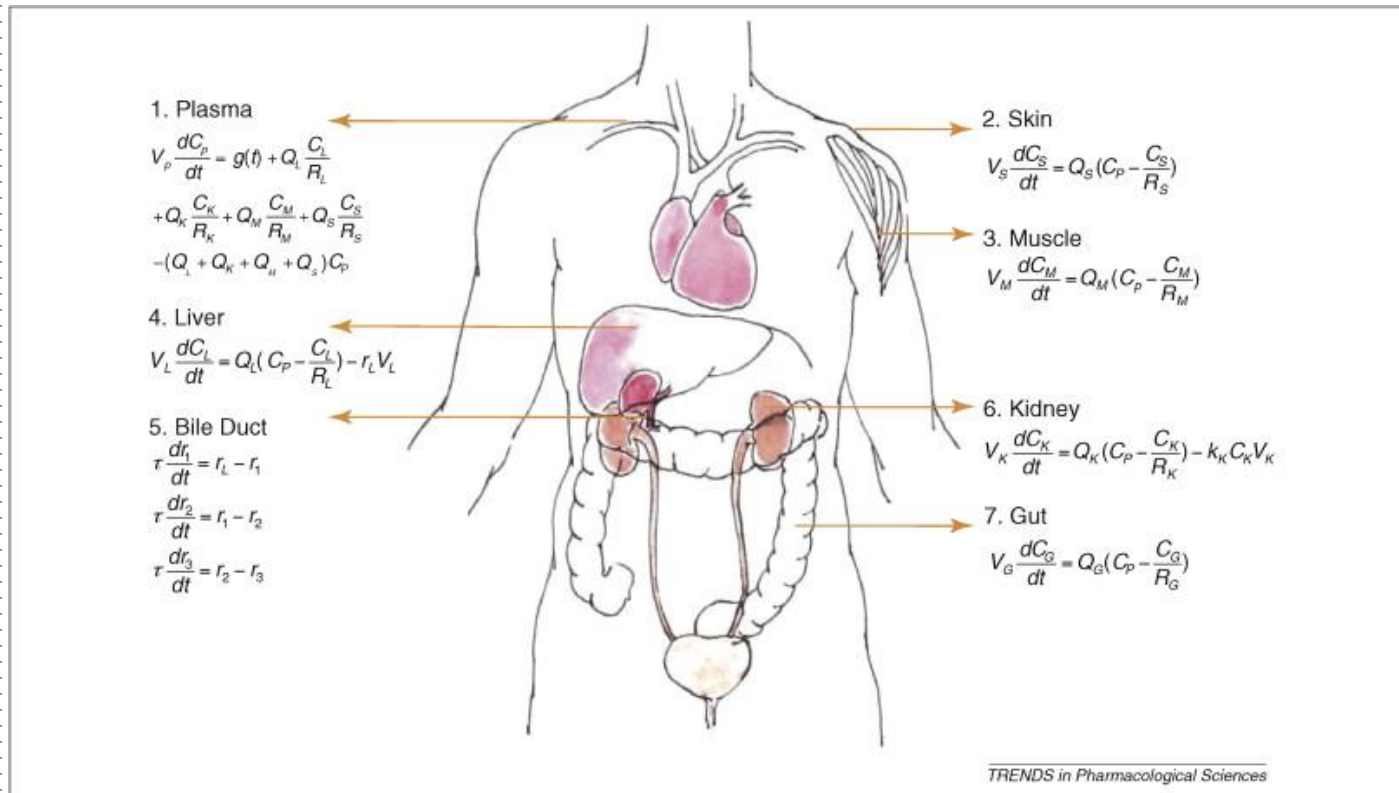
### Types of models:

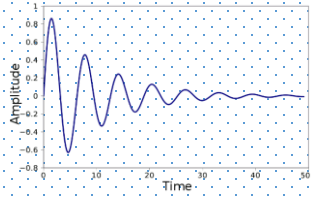
- Equation Based Models (EBM)
- Statistical Models (SM)
- System Dynamics Models (SDM)
- Agent Based Models (ABM)



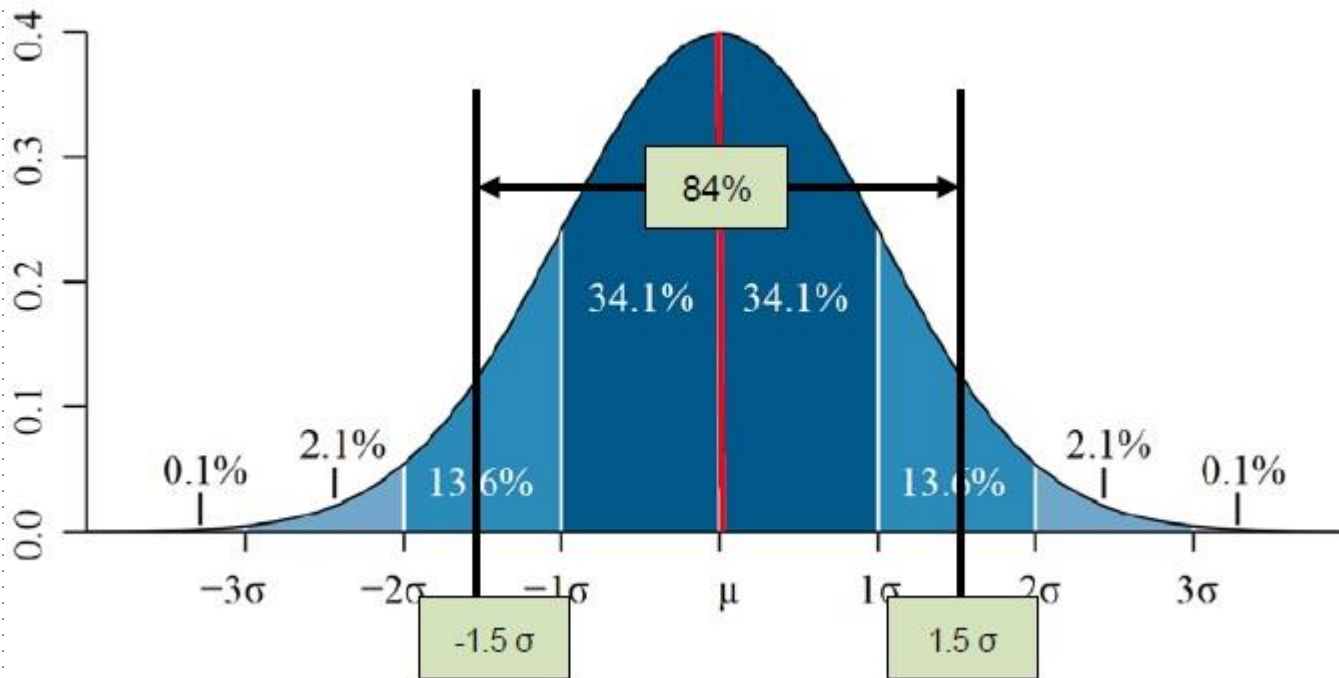


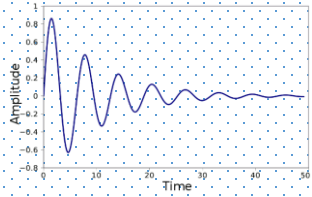
## Equation based Models (EBM)



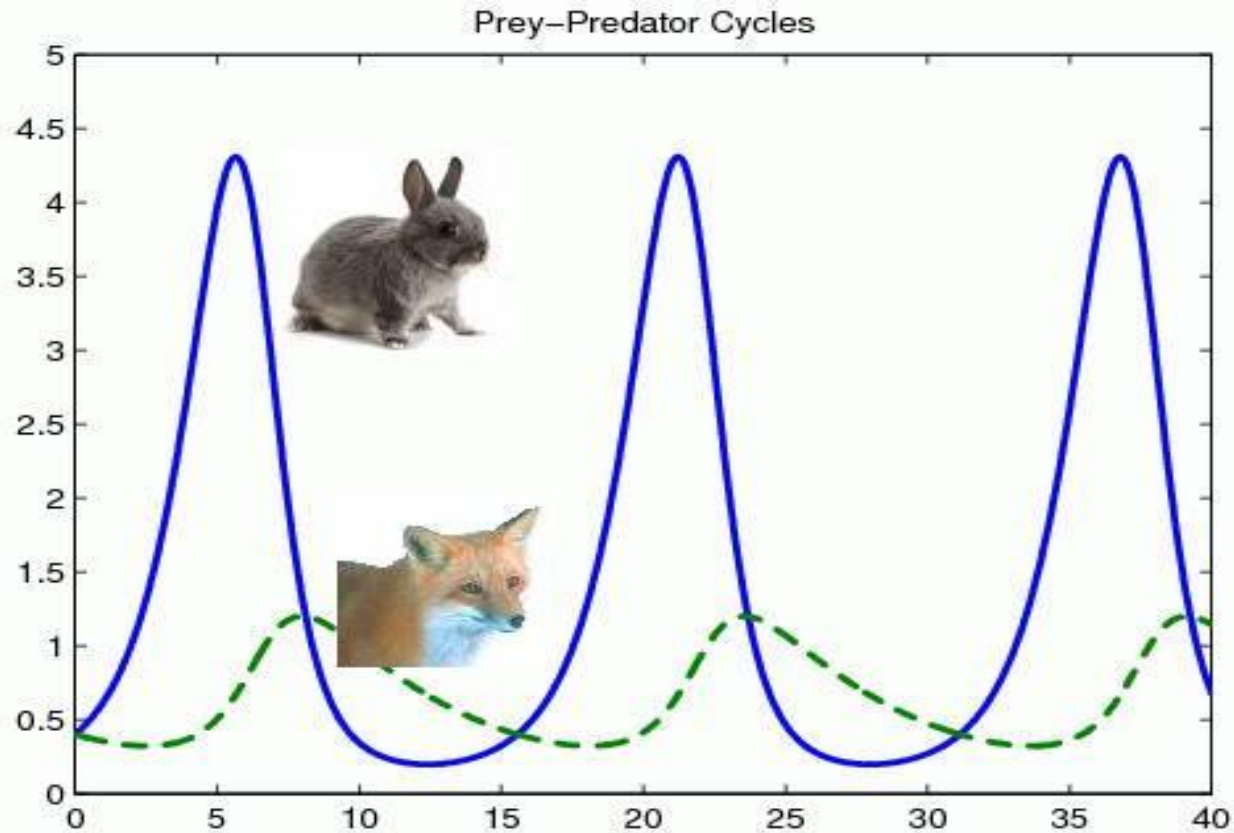


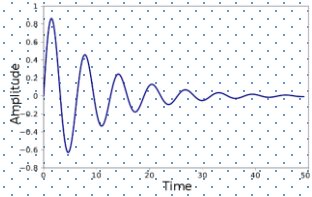
## Statistical Models (SM)





## System Dynamics Models (SDM)

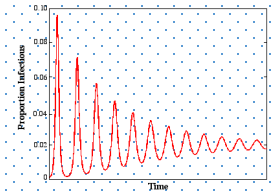




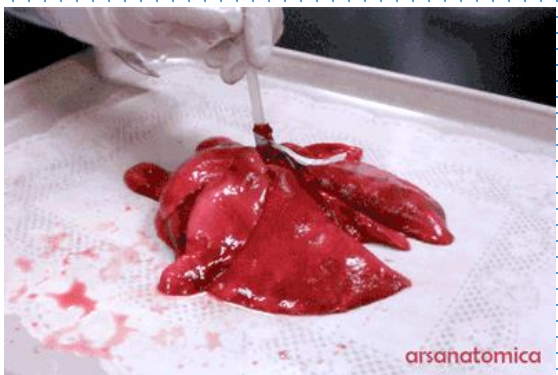
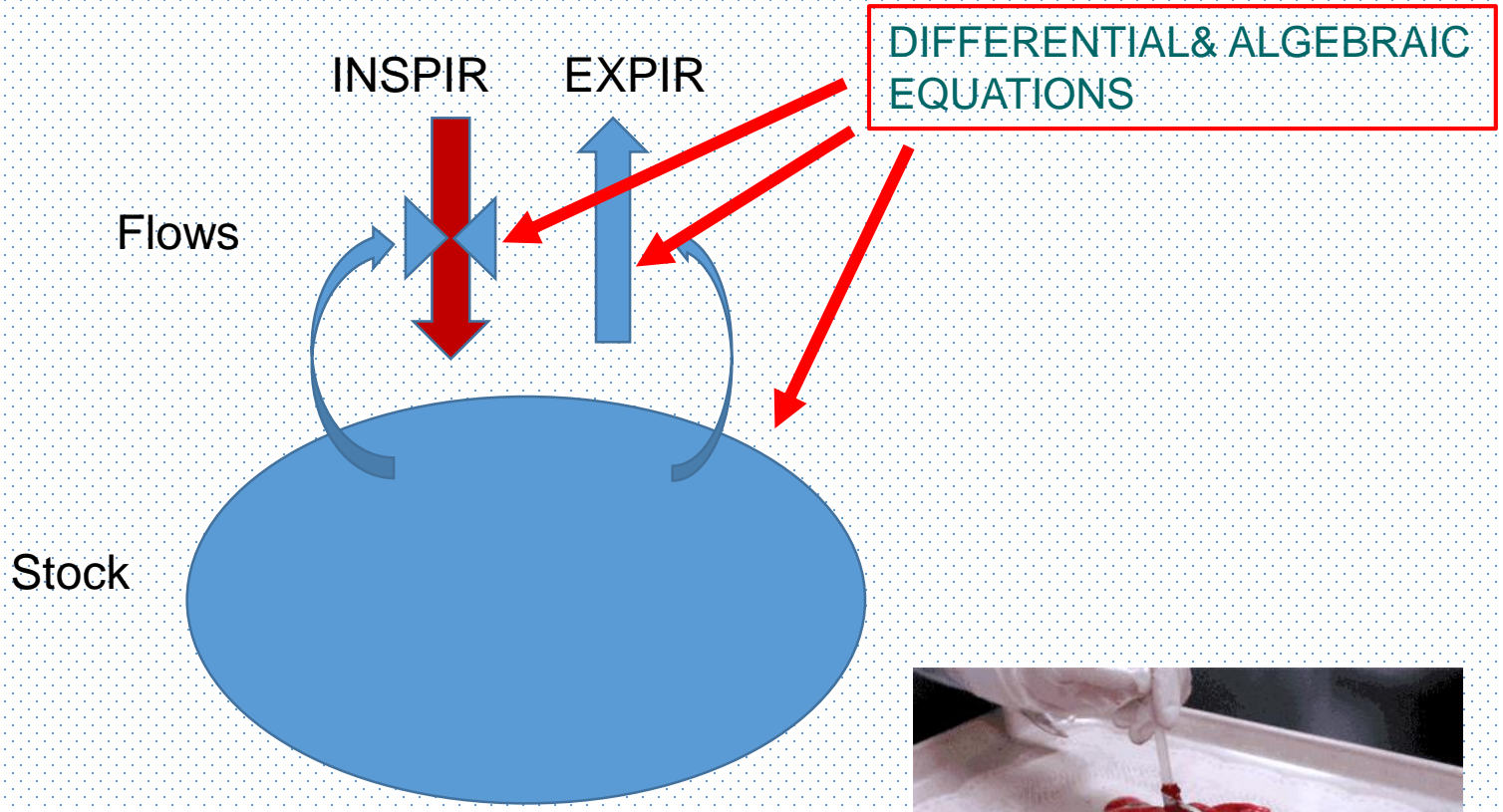
# Agent Based Models

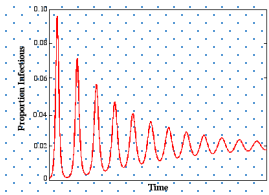
## Agent Based Modeling (ABM)



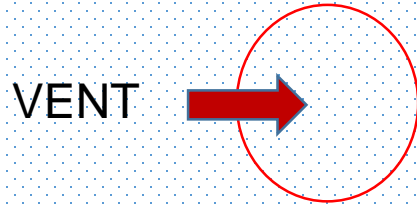


### BUILDING A MODEL



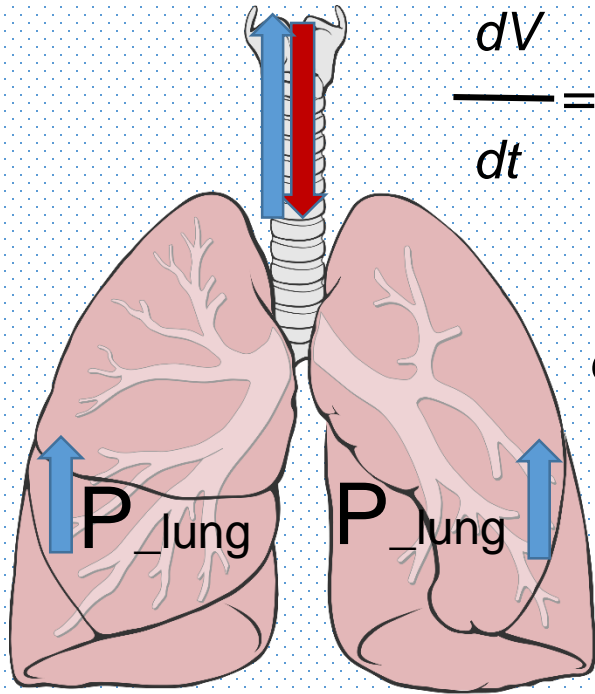


## BUILDING A MODEL

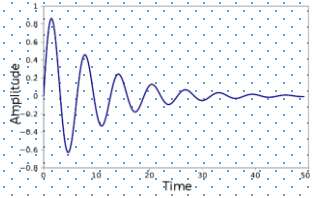


$$P_{mouth} = P_{atmos} + P_{drive}$$

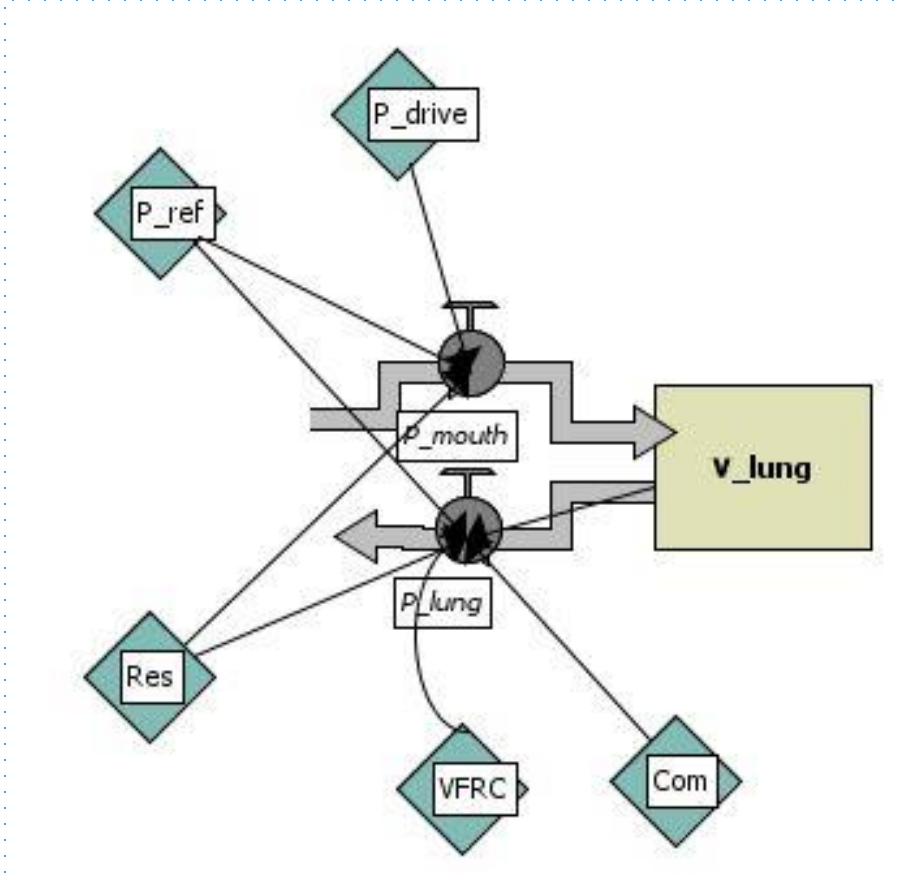
$$\frac{dV}{dt} = P_{mouth} - P_{lung} / Resist_{aw}$$

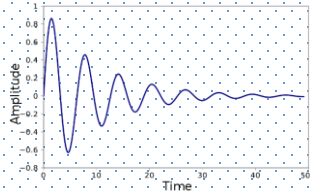


$$\frac{dP_{lung}}{dt} = P_{atmos} + V_{lung} - V_{FRC} / Compl$$

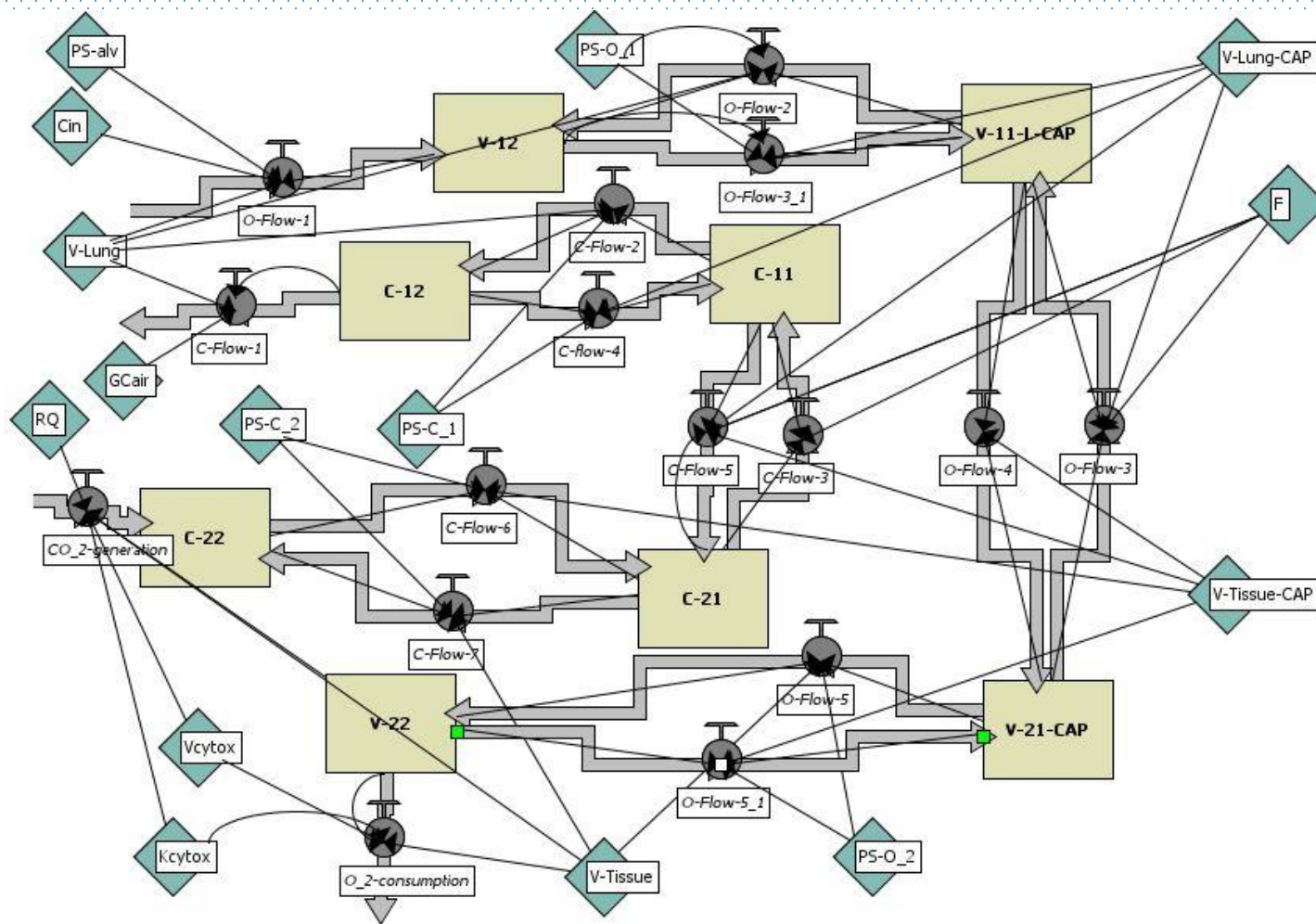


## BUILDING A MODEL: SDM scheme

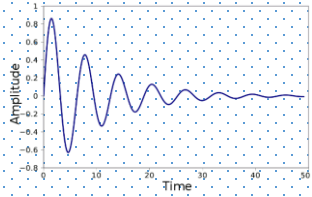




## O<sub>2</sub> & CO<sub>2</sub> transport & exchange: lung and peripheral tissues levels

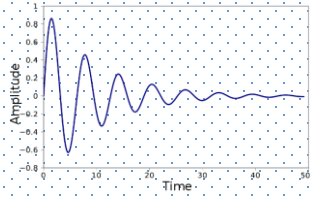






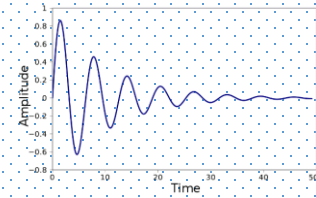
# ARTIFICIAL INTELLIGENCE (AI)





# ARTIFICIAL INTELLIGENCE (AI)





AI General

AI Narrow

## Artificial Intelligence (AI)

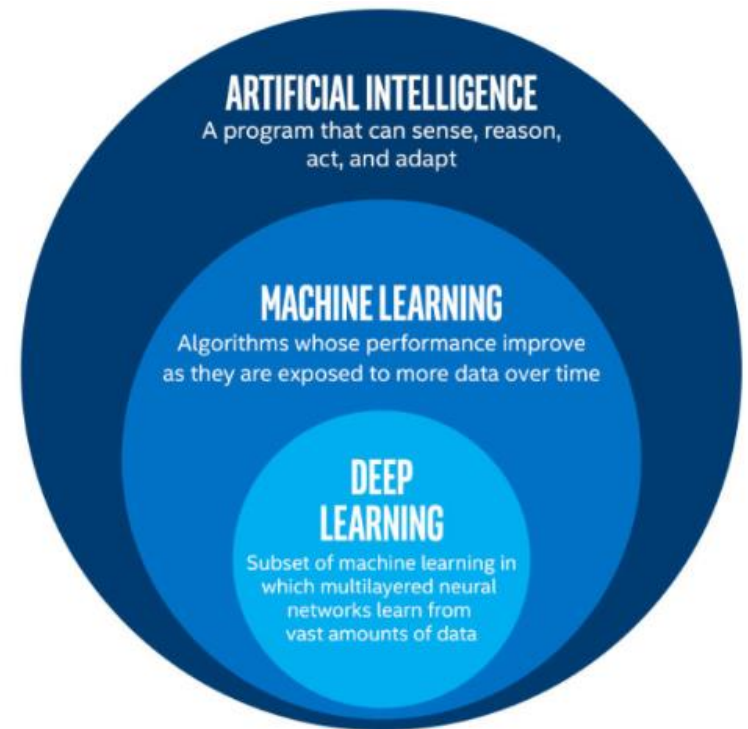
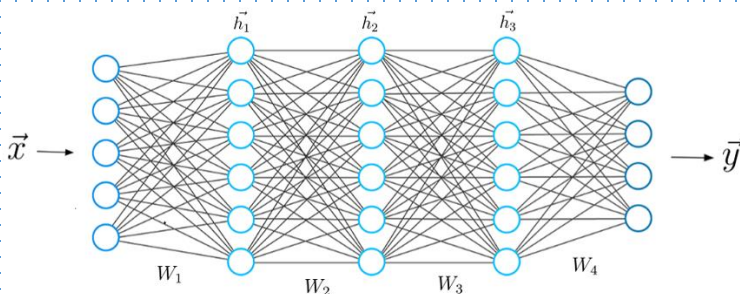
the possibility of computers to emulate human reasoning process

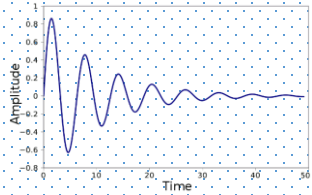
## Machine Learning (ML)

techniques / algorithms used by computers to learn / improve their performance in making informed decisions: supervised, unsupervised, reinforcement, transduction, multi-task, etc.

## Deep Learning (DL)

ANN based techniques:  
MLP, CNN, RNN





## AI - Examples

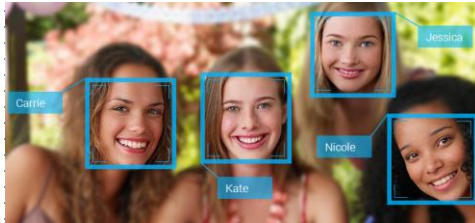


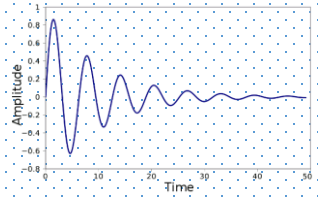
Image recognition  
(FB, Mento Park, CA)



Speech recognition  
(Apple's SIRI, Cupertino, CA)



Google, AlphaGo,  
G-apps  
(San Francisco, CA)



# AI in medicine / anesthesiology

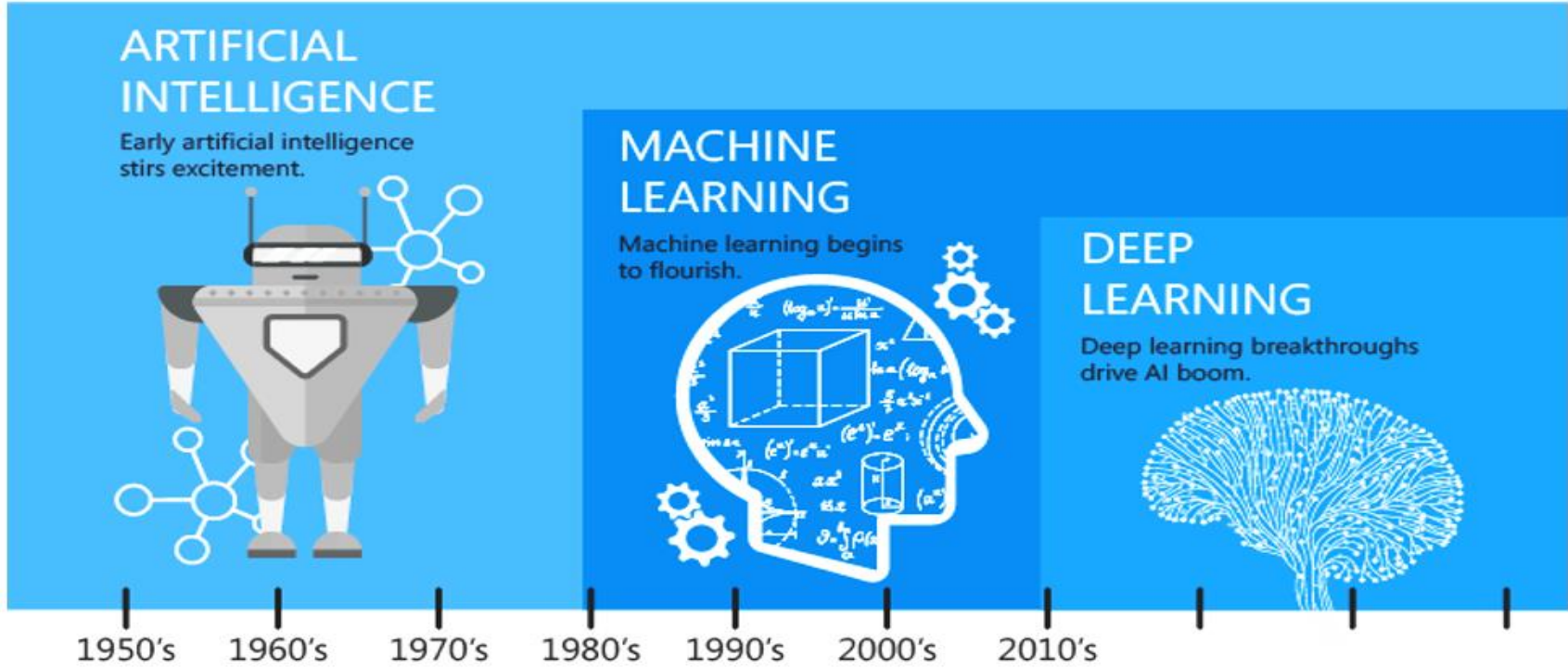
Bickford's Anesthesia System

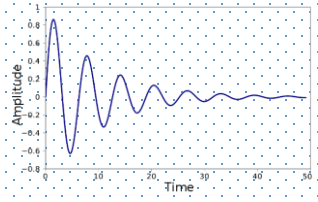
Rule-based Expert Systems

SedAsys, J&J, 2006  
McSleep, MUHC, 2008  
+ DaVinci MUHC, 2010

Kepler Intubation System, KIS teleanesthesia, 2011  
HumMod, 2011

SaMD, FDA, 2019



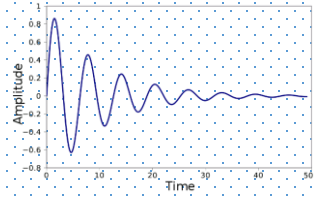


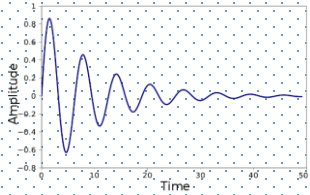
# AI: McSleepy



J

# Kepler Intubation System (KIS)

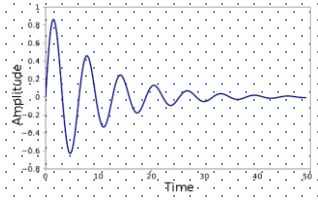




“Artificial Intelligence. It’s not intelligence, it might never be. The work of AI is at best clever use of algorithms to analyse large amounts of data. It has become a lazy term applied to too many tools and gadgets.”

*Andy Cotgreave, 2018*

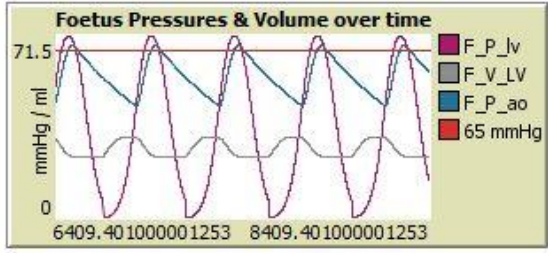
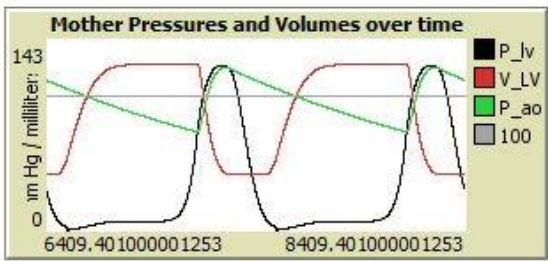




# Maternal-Foetal Circulation & Placenta / DO2 Model

setup go **Scenario**  
Test

Reset Test



SV\_LV ml/beat  
82.64

CO l/min  
7.44

EF %  
73

DO2 ml/min  
1175.99

DO2-Placenta  
104.75

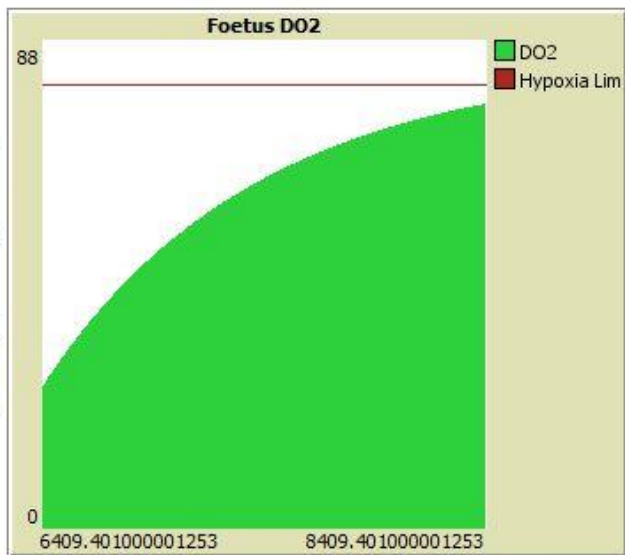
Placental O2/CO2 flow



Foetus SV\_LV ml...  
8.38

Foetus CO l/min  
1.13

Foetus DO2 ml/min  
76.42



Preload\_Volume-Status  
Normal

Myocardial-Contractility  
Normal-contractility

Afterload\_Vascular-Resistance-sys  
Normal-VR

Vascular-Elastance-sys  
Normal

Heart-Rate  
90

Hb\_m-slider  
120

Sa\_m\_O2-slider  
0.95

PS 0.2

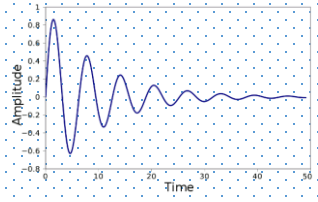
F\_Heart-Rate 135

Vol\_PL\_f-slider 0.14

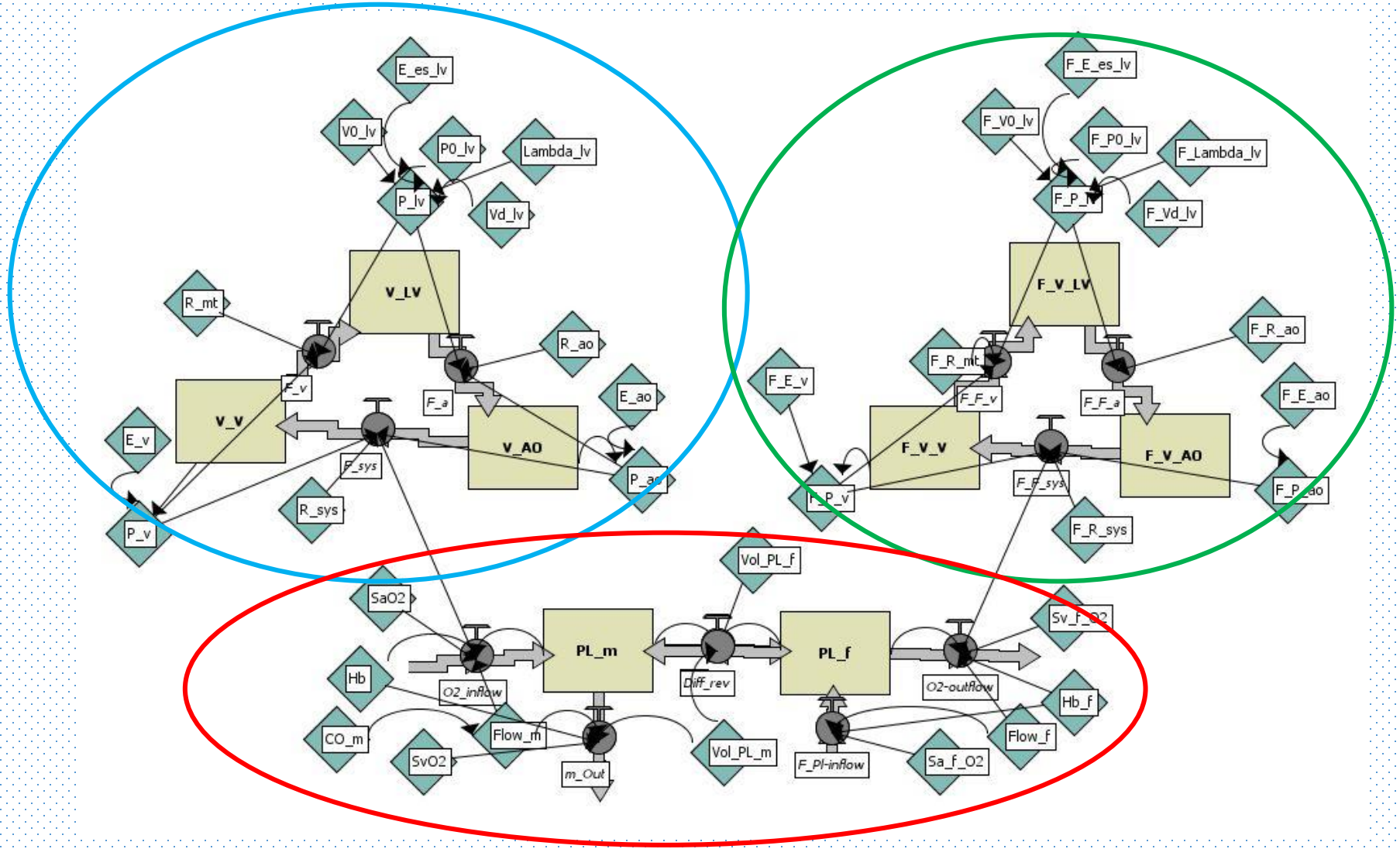
Hb\_f-slider 170

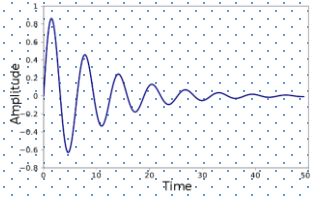
Vol\_PL\_m-slider 0.15

Sa\_f\_O2-slider 0.51

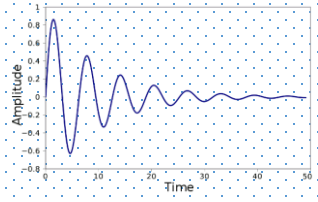


# Maternal-Foetal Circulation & Placenta / DO2 Model



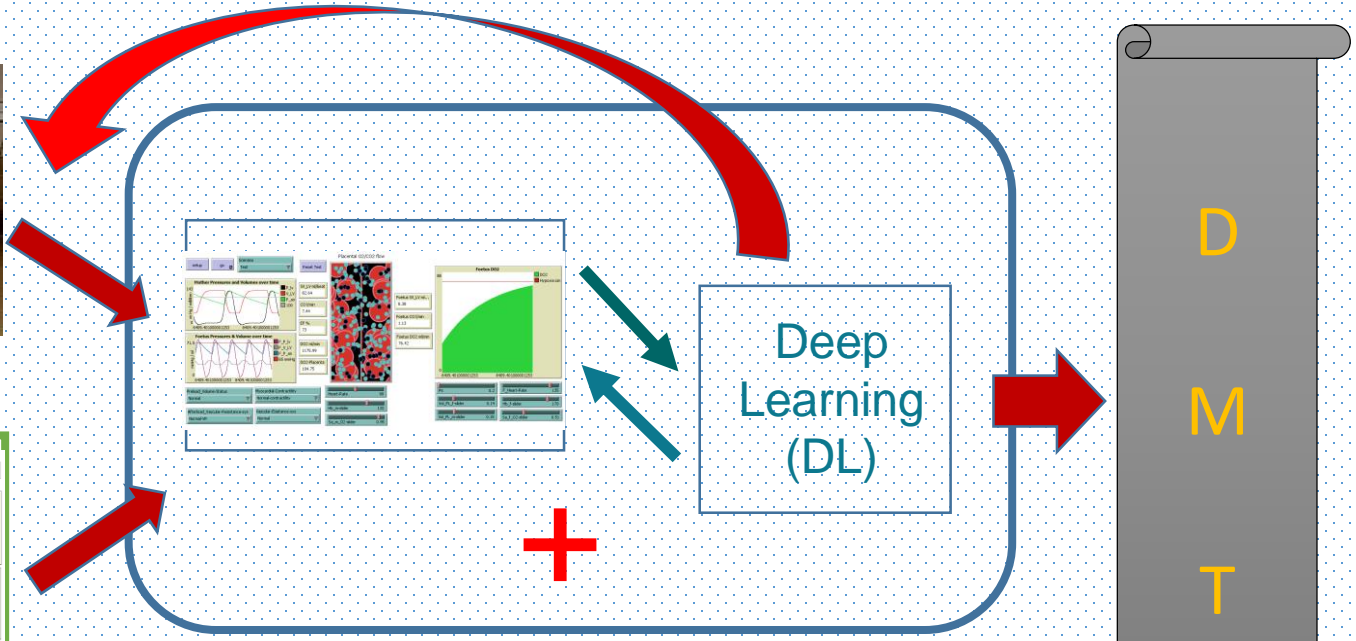
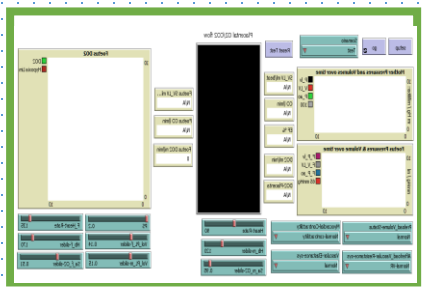


## Maternal-Foetal Circulation & Placenta / DO2 Model

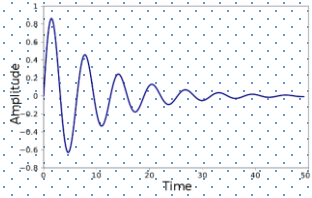


# INDIVIDUALISED MEDICINE

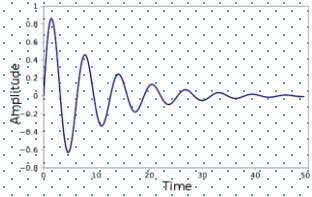
## Clinical Application of the Hybrid Model: Maternal-Foetal Circulation & Placenta / DO2



Multi-task learning  
(AI)



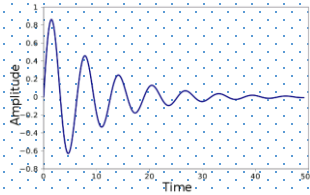
Future of AI ?



## SedAsys

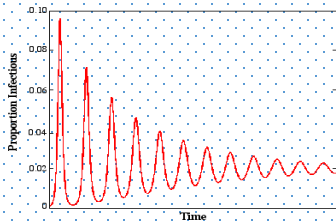


General anesthesia delivered by a computer system



“Recent innovations in artificial intelligence, especially machine learning, may usher in a new era of automation across many industries, including anesthesiology. It would be wise to consider the implications of such potential changes before they have been fully realized”

*John C. Alexander, MD, MBA, and Girish P. Joshi, MBBS, MD  
Department of Anesthesiology and Pain Management,  
University of Texas Southwestern Medical Center, Dallas, Texas  
2018*



- 01/ 2019 - 14 apps

- 07/ 2019 - 26 apps

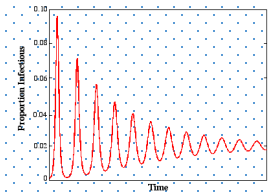
nowdays - > 30 apps

## Proposed Regulatory Framework for Modifications to Artificial Intelligence/Machine Learning (AI/ML)-Based Software as a Medical Device (SaMD)

*Discussion Paper and Request for Feedback*







## AI future: some possibilities

### ALGORITHMIC INFORMATION DYNAMICS (AID)



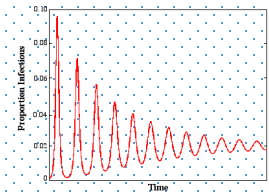
# 1 { H T H T H T H T H T }

# 2 { H T T T H H T H H T }

Shannon's Entropy(statistics)

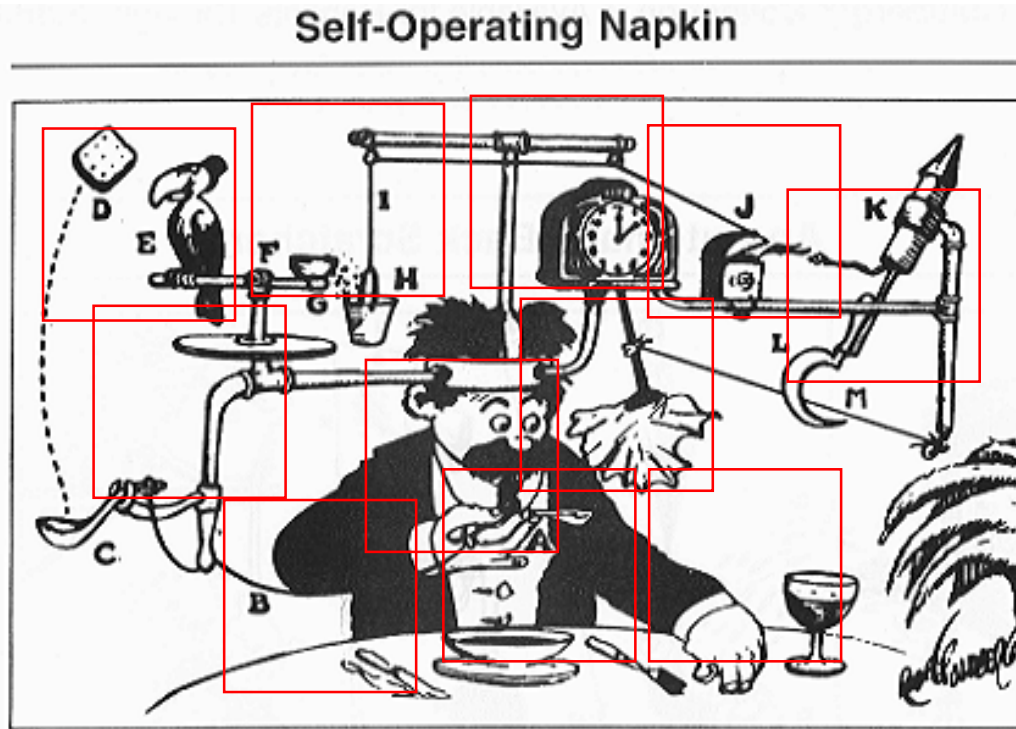
≠

AID



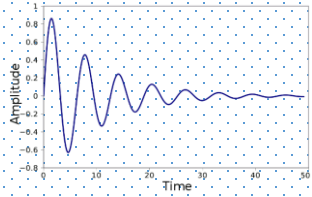
## AI future: some possibilities

### ALGORITHMIC INFORMATION DYNAMICS (AID)



BLOCK DECOMPOSITION METHOD -  $\Sigma$  complexity

+ Deep Learning



<http://modelingcommons.org/account/models/2495>  
[https://www.complex-systems.com/abstracts/v28\\_i01\\_a03/](https://www.complex-systems.com/abstracts/v28_i01_a03/)  
[https://www.researchgate.net/profile/Victor\\_lapascurta/research](https://www.researchgate.net/profile/Victor_lapascurta/research)