

AI Application in Health

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Sumary

The Evolution of Machines

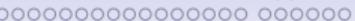
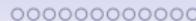
Data Analytics in Health

Application to Physiological Signals: Arousal detection

Application to EHR

Application to EHR + Physiological Signals

Practical Information on the Course



Machines are beating us at everything

"They, the machines, will make decisions for us, they will govern us."

Ethics for Machines, Jose Ignacio Latorre

- Initially, machines replaced us in physical labor.
 - Then in calculation.
 - Now there are machines making decisions that are less and less trivial.
 - In the future, machines will increasingly make ethical decisions.

And we like to **delegate decisions**.

How do Artificial Intelligences Learn?

Two strategies:

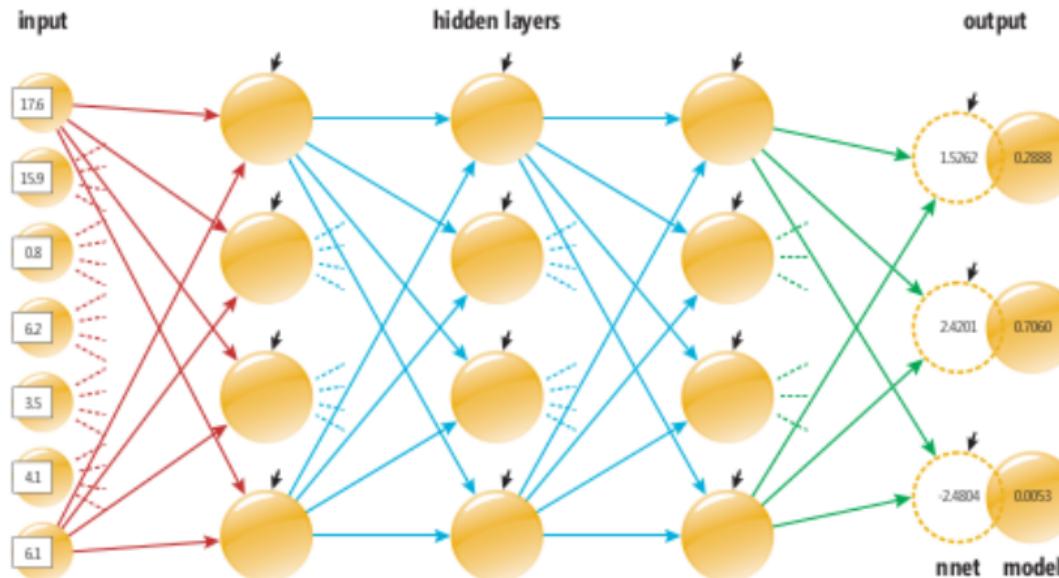
Expert systems (80's):

- Humans program their knowledge into detailed algorithms.
 - Advanced statistics.
 - Bayesian inference.
 - Ex: Medical diagnosis.
 - It was a **failure**.
 - **AI winter**.

Neural networks:

- Learning from examples.
 - Copy the human brain, not its logic.
 - Learning = Training.

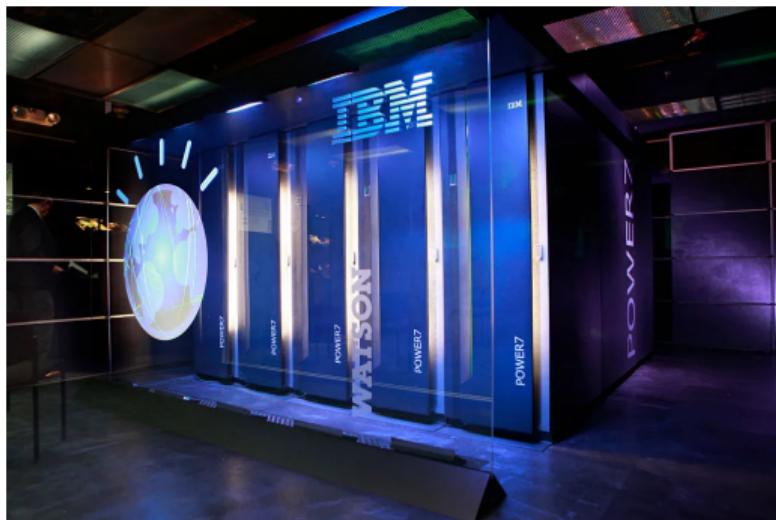
Neural networks learn from examples



Great successes: AlphaGo

Medical Assistant: IBM Watson

Delegation of Decisions



- Doctors can now have a virtual assistant.
- Trained with thousands of cases.
- Thousands of treatments.
- Detailed casuistry. /item Inference from inaccurate data.

Is this the way?

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What are the main problems in Healthcare Systems? (with people living 100 years)

In developed countries

The healthcare expenditures are growing in an unsustainable way

In underdeveloped countries

There are not enough resources for taking care of a growing population

How can AI help to solve this problems?

Areas where AI can Help

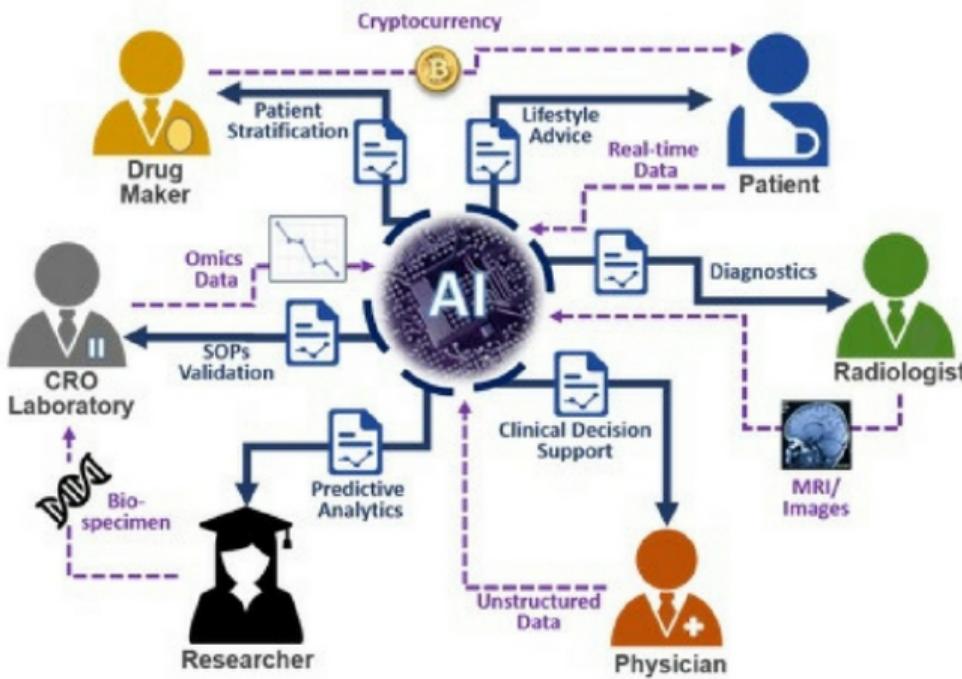
AI can help to reduce costs in two ways (at least)

1. Clinical Decision Making

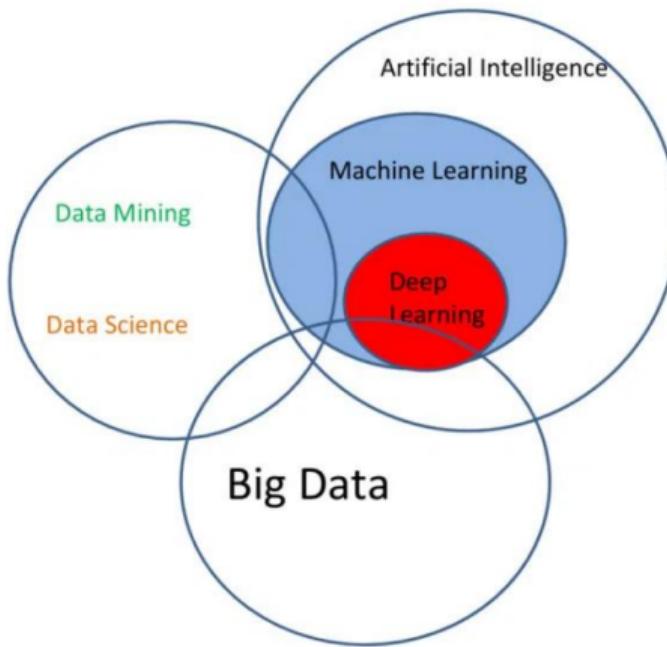
- Avoiding misdiagnosis and unnecessary treatments
 - Personal recommend treatments
 - Preventive medicine. Models for risk prediction

2. Operational planning. Resource allocation

Artificial Intelligence and Medicine



Differences between AI, Machine learning and Data Analytics

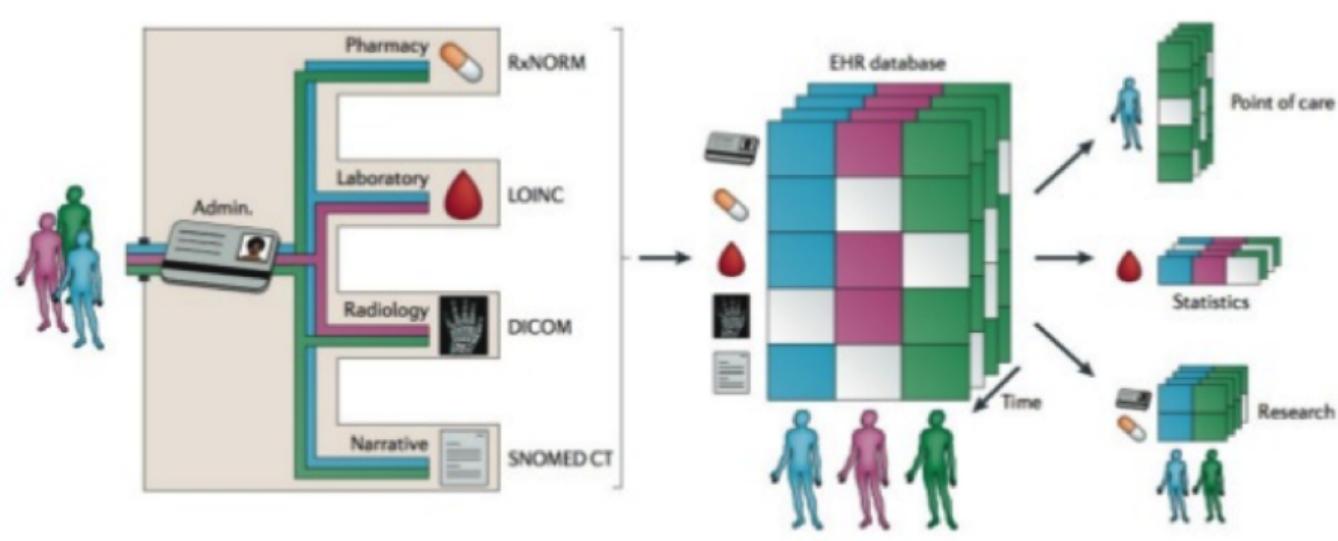


Why Deep Learning Now?

From 2012

- Big Datasets
 - Images
 - Vital signals/Sensors (ICU, wearables, ...)
 - EHR
 - Laboratory Data
 - Genome sequences
 - Events sequences
- GPU availability (High Performance Computing)
- Cloud Computing
- Open source algorithms (Tensorflow, Pandas, SkLearn, ...)

Medical Data & Electronic Health Record (EHR)



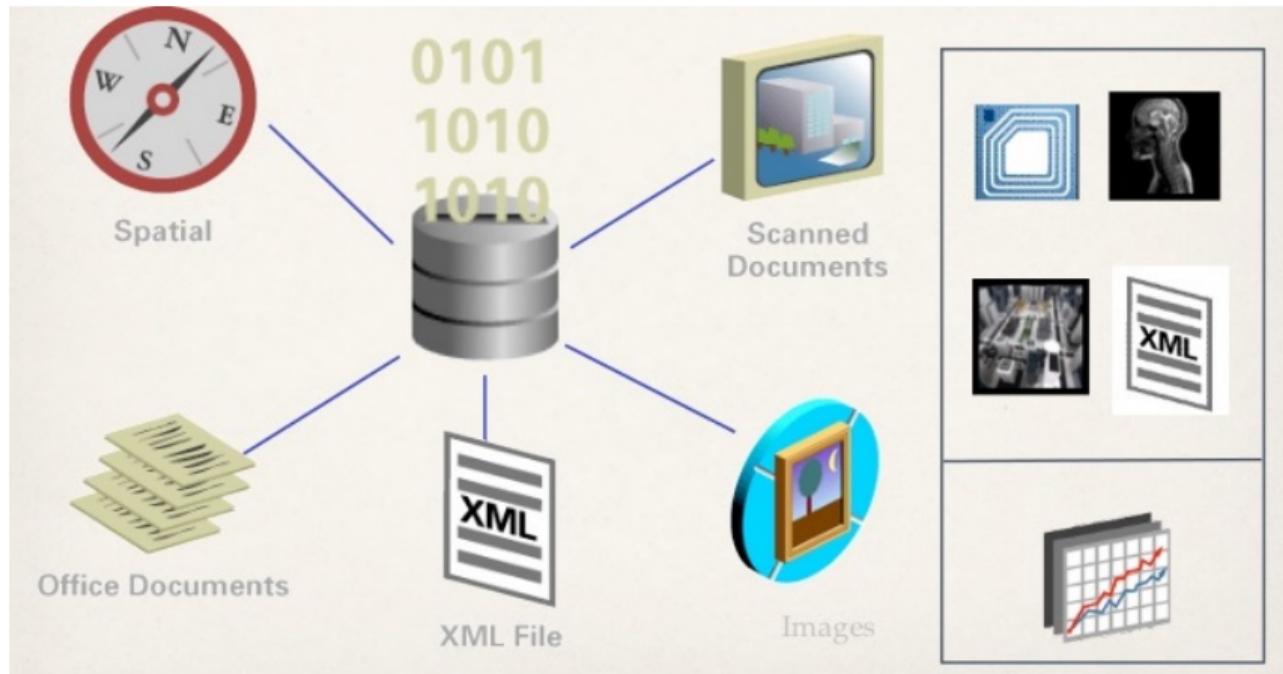
What Kind of Clinical Data Can Be Found in a Hospital?

Structured Data - The Electronic Health Register (EHR)

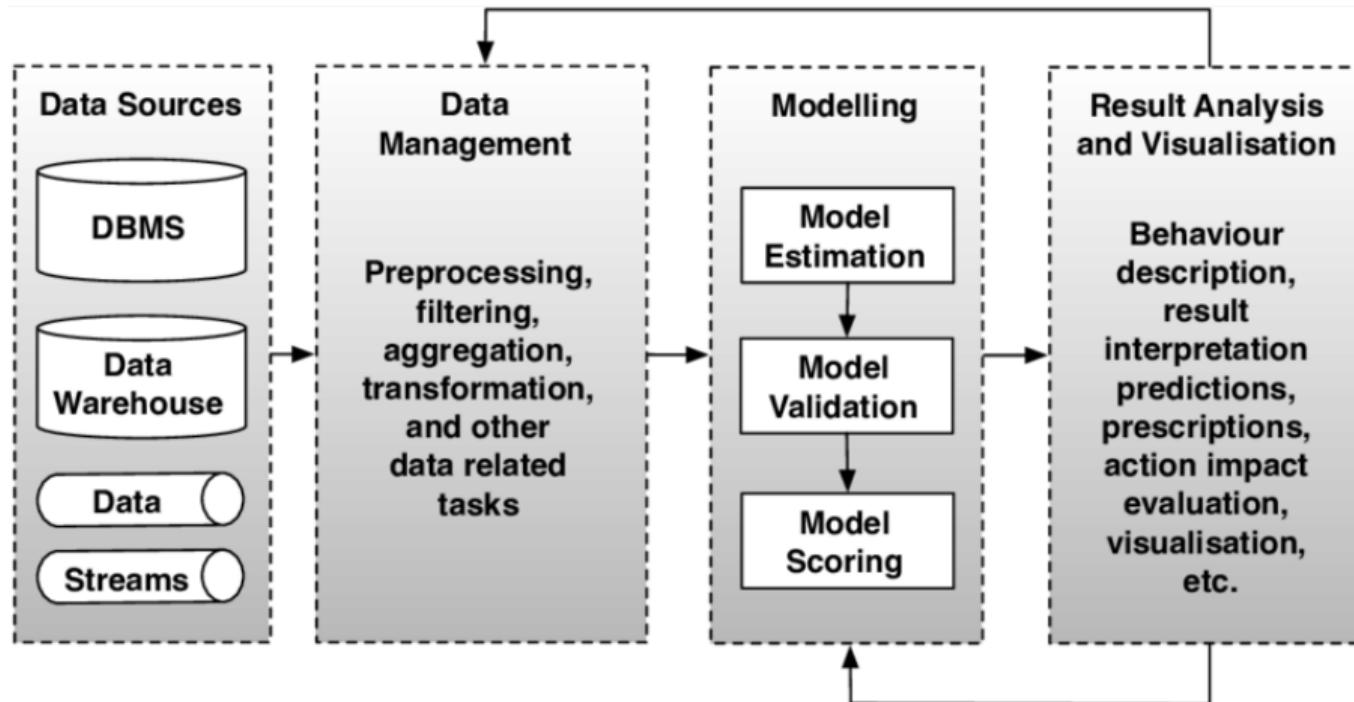
	Assigned diagnosis				Medications			Laboratory values			Demographics	
	C1	C2	C3	C4	M1	M2	M3	L1	L2	L3	D1	D2
Patient 1	■	■						■				
Patient 2												
Patient 3			■									
Patient 4	■							■				
Patient 5			■		■							
Patient 6		■										
Patient 7			■		■							
Patient 8			■			■						
Patient 9	■		■					■	■			

What Kind of Clinical Data Can Be Found in a Hospital?

Unstructured Data: Images, Clinical reports, Time Series

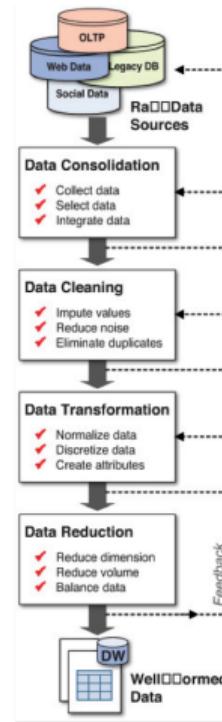


Typical Workflow in Data Analytics

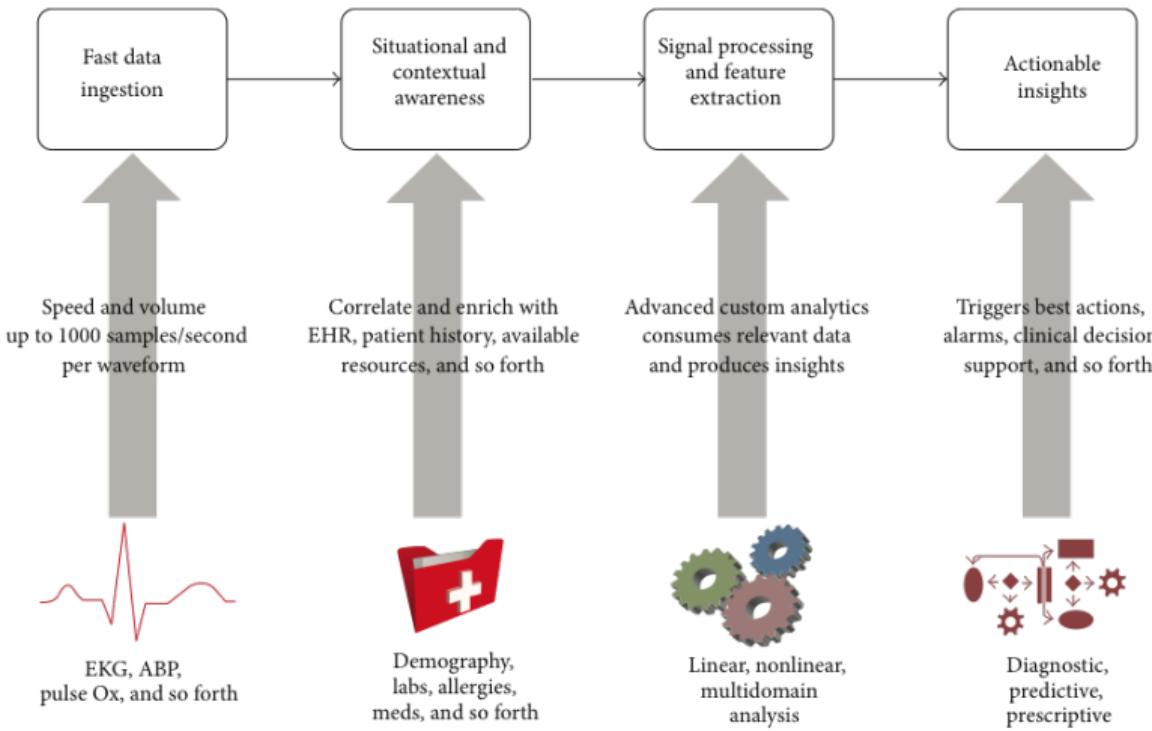


Data Preprocessing

- Time consuming
- Very important
- Anonymization is a previous step to preprocessing



Big data Processing



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Application to Physiological Signals: Arousal detection

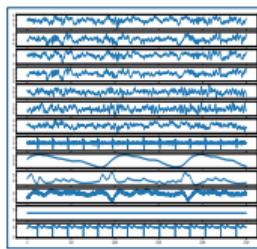
Knowledge extraction based on wavelets and DNN for physiological signals (PS): Arousal case

- **Goal:** Detection of arousal zones when sleep
- Useful to detect sleep disorders
- Sleep (-4-9 hours) = ... non-REM + REM + non-REM + ... (REM: Rapid Eye Movement)
- Signal we analyze: ECG + VOmax + EEG + EOM
- Non-balanced data
- Medical collaboration is needed.

Application to Physiological Signals: Arousal detection

Knowledge extraction based on wavelets and DNN for physiological signals (PS): Arousal case

1. Slide windows, of 13 PS

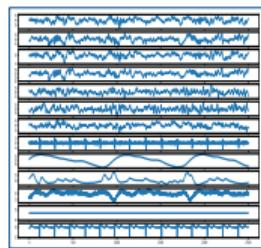


Medical decision support based on discrete wavelet transform for feature extraction and classification of 13 PS, >1850 patients

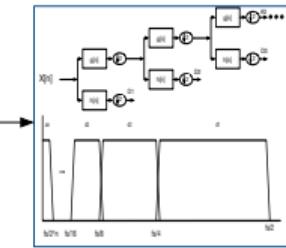
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2. DWT: Details extraction

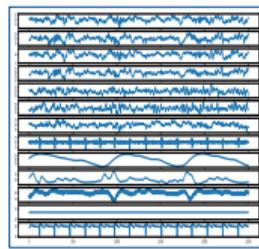


Medical decision support based on discrete wavelet transform for feature extraction and classification of 13 PS, >1850 patients, 6M registers.

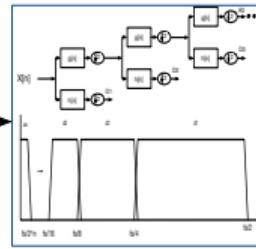
Application to Physiological Signals: Arousal detection

Knowledge extraction based on wavelets and DNN for physiological signals (PS): Arousal case

1. Slide windows, of 13 PS



2. DWT: Details extraction



3. Feature extraction from details

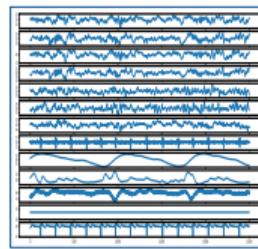
- Mean of absolute values
- Average power
- Standard deviation
- Ratio of the absolute mean values of adjacent sub-bands.

Medical decision support based on discrete wavelet transform for feature extraction and classification of 13 PS, >1850 patients, 6M registers. Information reduction from 32500 samples per patient to 312

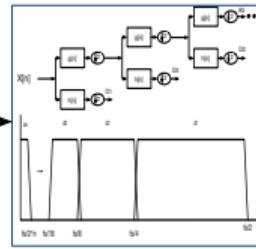
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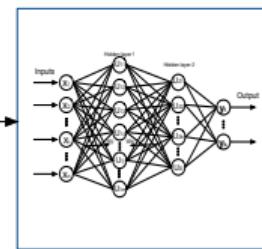
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4. DNN for classification

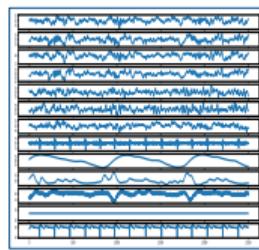


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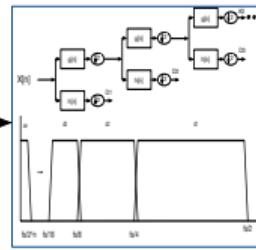
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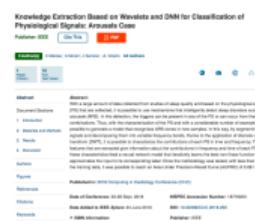
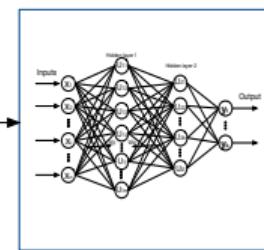
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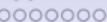
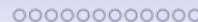
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Practical Information on the Course



Case 1: Mortality Prediction for AKI

Acute Kidney Injury (AKI)



Affects kidney **structure**

and **function**

30 - 70 % Intensive Care
Unit (ICU) incidence



Case 1: Mortality Prediction for AKI

Acute Kidney Injury (AKI)



Affects kidney structure

and function

Exposure to critical

situations

30 - 70 % Intensive Care Unit (ICU) incidence

- Sepsis
 - Trauma
 - Cardiac surgery



Case 1: Mortality Prediction for AKI

Acute Kidney Injury (AKI)



Affects kidney structure
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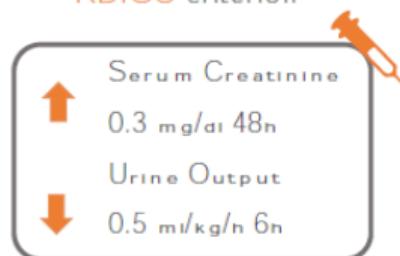
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Exposure to critical
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KDIGO criterion



Case 1: Mortality Prediction for AKI

Acute Kidney Injury (AKI)



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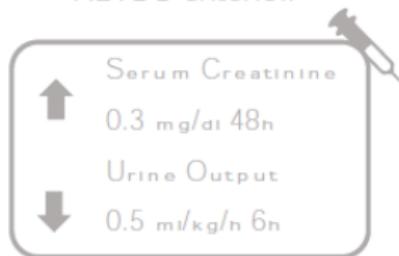
30 - 70 % Intensive Care
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Exposure to critical
situations

- Sepsis
- Trauma
- Cardiac surgery

KDIGO criterion



MORTALITY

5 % of all hospitalized
patients

10 - 25 % of Intensive
Care Unit (ICU) patients

*Jaber, B. L. et al. World Incidence of AKI: A Meta-Analysis. *Clin. J. Am. Soc. Nephrol.* 8, 1482–1493 (2013). 2

Case 1: Mortality Prediction for AKI

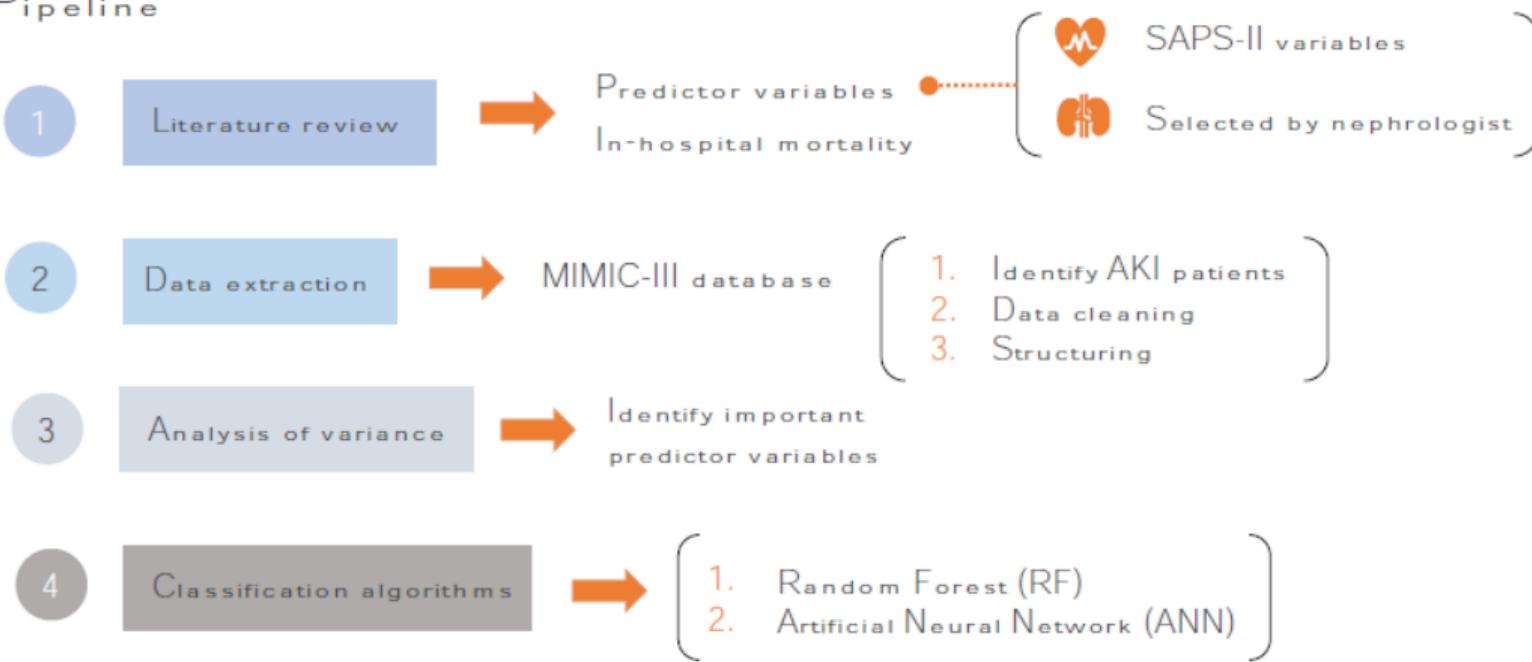
Objectives

- General **Predict** mortality in-hospital in patients with AKI in the ICU
- Secondary Select and analyze the most important predictors variables (**Knowledge Discovery**)

Case 1: Mortality Prediction for AKI

Methods

Pipeline



Methods

Variables and Metrics

Variables

Factors	SAPS-II	Selected by an expert
Age Type of admission Relevant diseases Gender Number of admissions	Heart Rate	Creatinine
	Systolic Blood Pressure	Glucose
	Temperature	Platelet Count
	GCS	Red Blood Cells
	P _a O ₂	Chloride
	F _i O ₂	Anion Gap
	Urine Output	Magnesium
	BUN	PT
	Sodium	PTT
	Potassium	Phosphate
	Bicarbonate	Calcium
	Bilirubin	pH
	WBC	pCO ₂
		pO ₂
		Lactate
		Lymphocytes
		Eosinophils
		Neutrophils
		Alanine Aminotransferase
		Aspartate Aminotransferase
		Alkaline Phosphatase

Metrics

		Predicted	
		Alive	Dead
Confusion Matrix	Actual	Alive	TN
		Dead	FN
			TP

General performance:

- Area Under Receiving Operation

Characteristic (AUC)

$$\text{Accuracy} = \frac{\text{TP}+\text{TN}}{\text{TP}+\text{TN}+\text{FP}+\text{FN}}$$

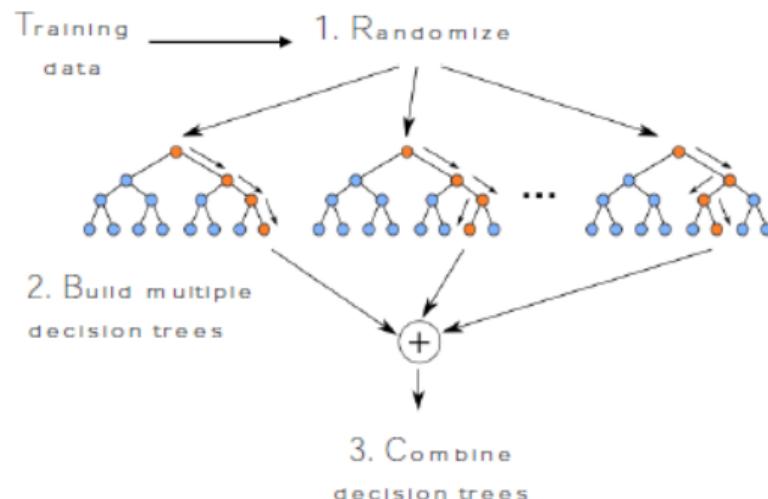
⎛ Biased parameter for
 unbalanced groups ⎝

Classification Algorithms

Random Forest

Some features

- RF is very easy to set up. Good algorithm for starting
- Used as baseline system
- Even with small data works properly
- Rate the importance of the attributes

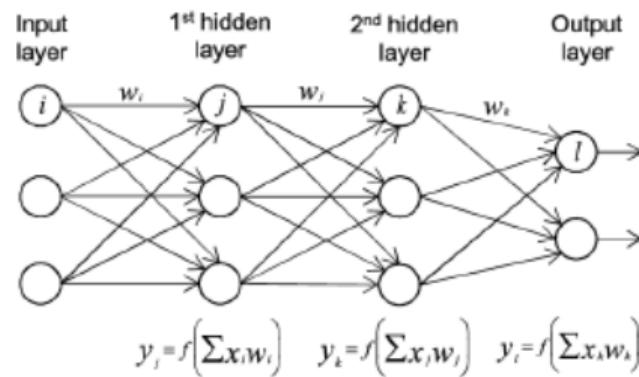


Classification Algorithms

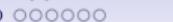
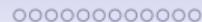
Artificial Neural Networks

Some features

- Usually gets the best performance
- Lots of parameter to tune, difficult
- With small data tends to overfit or do not get a good performance



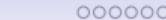
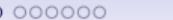
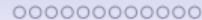
Training with validation: 85% data
Test evaluation: 15% data



Results

Comparison performance: Clinically valuable metrics

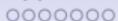
	Sensitivity		NPV		AUROC	
	RF	ANN	RF	ANN	RF	ANN
1. SAPS-II 4153	63.7%	68.8%	78.7%	80.3%	0.79	0.77
2. SAPS-II stat. 6754	89.3%	81.5%	92.4%	89.0%	0.77	0.76
3. SAPS-II stat. 15435	74.7%	77.7%	89.9%	90.6%	0.82	0.82
4. SAPS-II 15435	87.1%	85.7%	94.2%	93.8%	0.87	0.87
5. SAPS-II + analytics 15435	76.5%	79.9%	91.7%	92.5%	0.85	0.85



Results

Comparison performance: Other metrics

	Specificity		Precision		Accuracy	
	RF	ANN	RF	ANN	RF	ANN
1. SAPS-II 4153	80.7%	76.6%	66.5%	63.9%	74.3%	73.6%
2. SAPS-II stat. 6754	49.9%	57.2%	40.6%	42.2%	60.8%	63.9%
3. SAPS-II stat. 15435	74.1%	71.1%	48.8%	46.9%	74.3%	72.7%
4. SAPS-II 15435	69.8%	71.4%	48.7%	49.7%	74.1%	75.0%
5. SAPS-II + analytics 15435	79.6%	75.9%	53.5%	50.5%	78.9%	76.9%



Discussion

Strengths and Drawbacks

Strengths

- Good performance in relevant metrics
- High potentials on clinical applications
- Use of a public-available and large-scale database

Drawbacks

- Insufficient results on metrics related to resource management
- Retrospective study
- Not able to implement yet in clinics

Case 1: Mortality Prediction for AKI

Conclusions

Predictive model of in-hospital mortality for AKI patients in the ICU

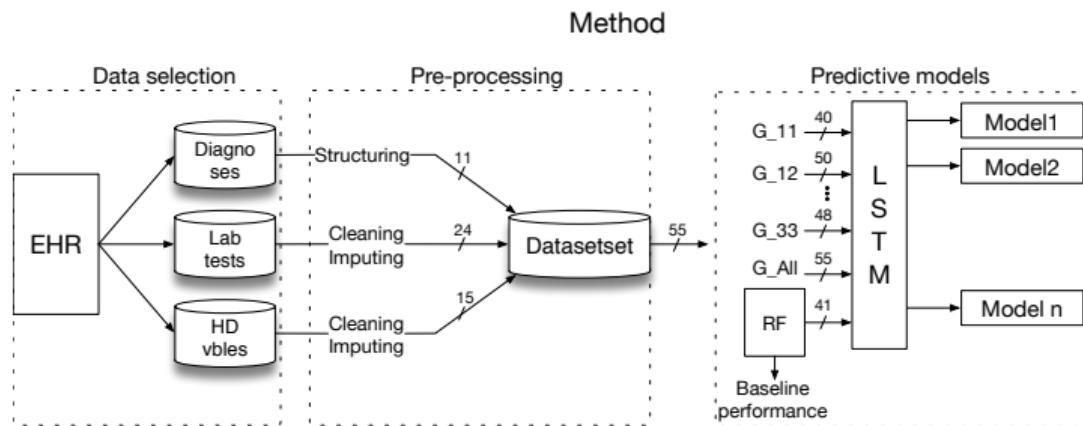
- Increase on the number of subjects
- Increase on the number of predictor variables

High potential in clinical practice

- Take decisions in critical situations
- Redistribute/allocate resources in ICU
(Ethical Issues)

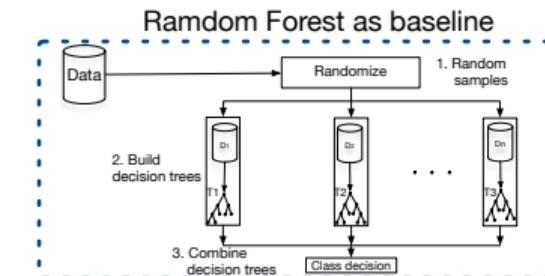
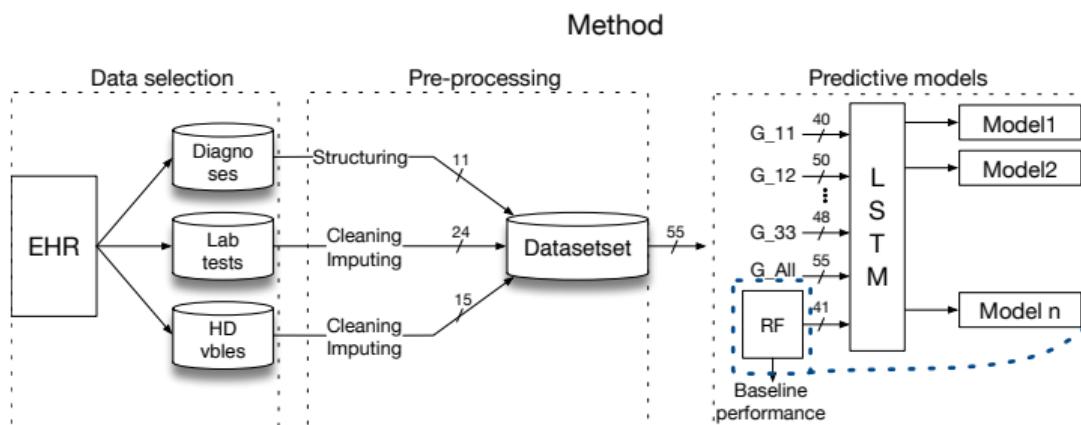
Case 2: Mortality Prediction in ESRD

Mortality prediction enhancement in end-stage renal disease: A machine learning approach



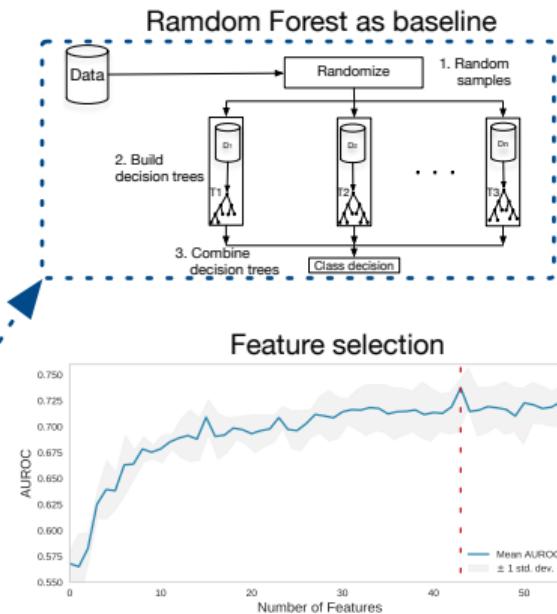
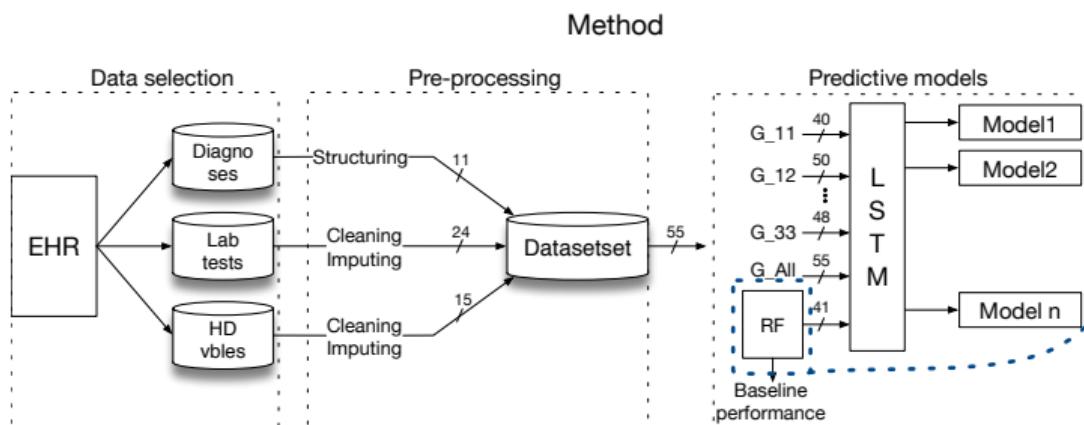
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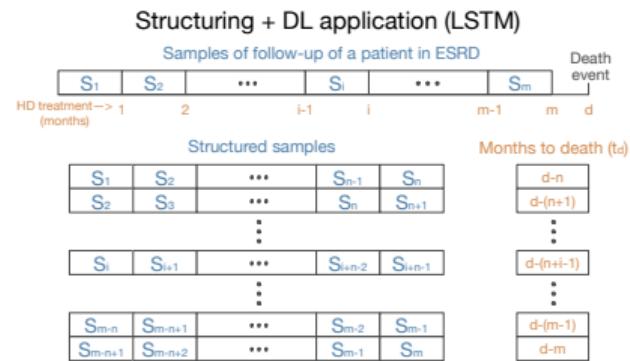
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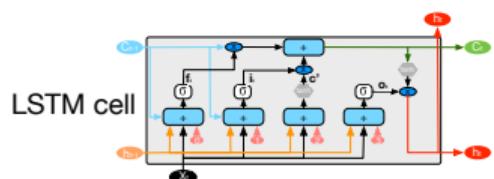
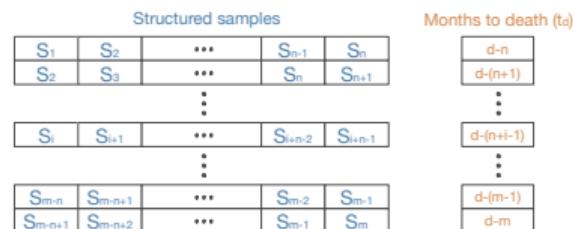
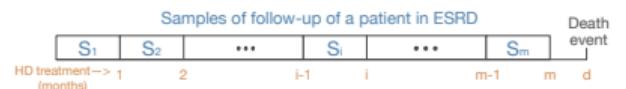
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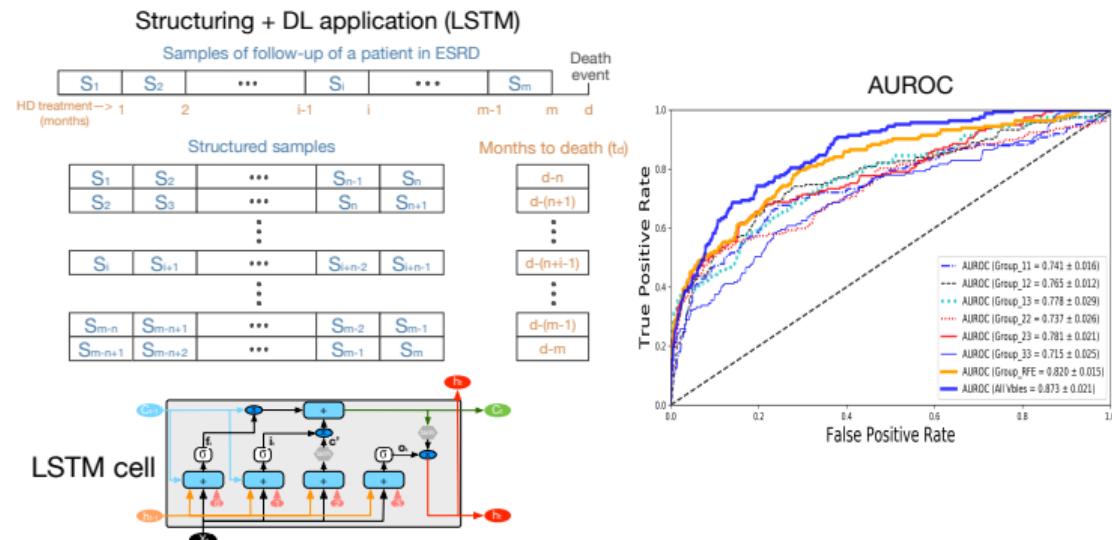
Mortality prediction enhancement in end-stage renal disease: A machine learning approach

Structuring + DL application (LSTM)



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Structuring + DL application (LSTM)



Structured samples

S_1	S_2	...	S_{n-1}	S_n
S_2	S_3	...	S_n	S_{n+1}

S_1	S_{i+1}	...	S_{n-2}	S_{n+1}
S_i	S_{i+2}	...	S_{n-1}	S_n

S_{m-n}	S_{m-n+1}	...	S_{m-2}	S_{m-1}
S_{m-n+1}	S_{m-n+2}	...	S_{m-1}	S_m

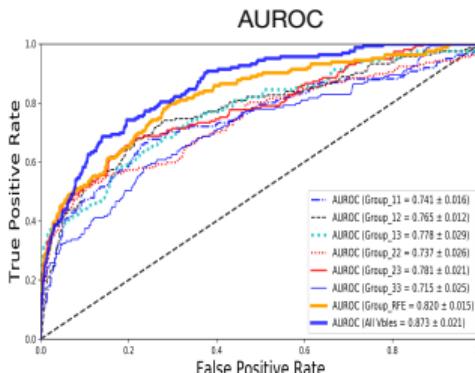
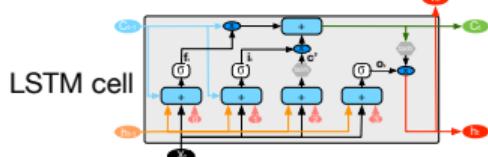
Months to death (t_d)

$d-n$
$d-(n+1)$

$d-(n+1)$
$d-(n+2)$

$d-(n+2)$
$d-(n+3)$

$d-(n+3)$
$d-m$



Mortality prediction enhancement in end-stage renal disease: A machine learning approach

Fabio Mazzoni, Antonio Mazzoni, Gianni Sestini, Jean-Luc Vincent, Jean-Baptiste Mazzoni

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Informatics in Medicine Unlocked

Volume 1, Number 1, March 2018

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Highlights

- Machine learning approaches can lead to a paradigm shift in the analysis of predictive factors for mortality in ESRD.
- The massive use of variables together with artificial neural networks improves predictive models of mortality in ESRD.
- Machine learning approaches can reveal causal relationships in variable not explored before by the expert staff.

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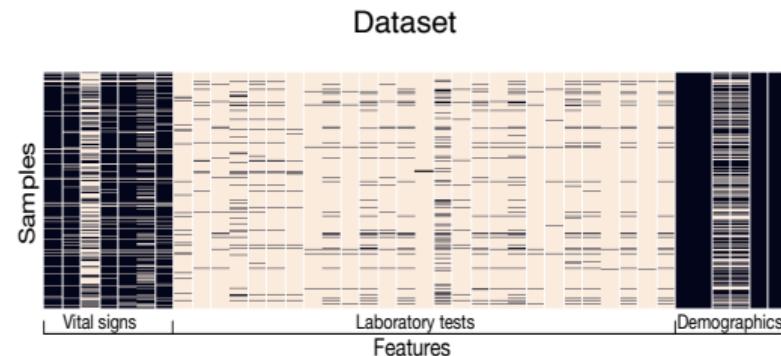
Application to EHR + Physiological Signals

Practical Information on the Course



Application to EHR + Physiological Signals

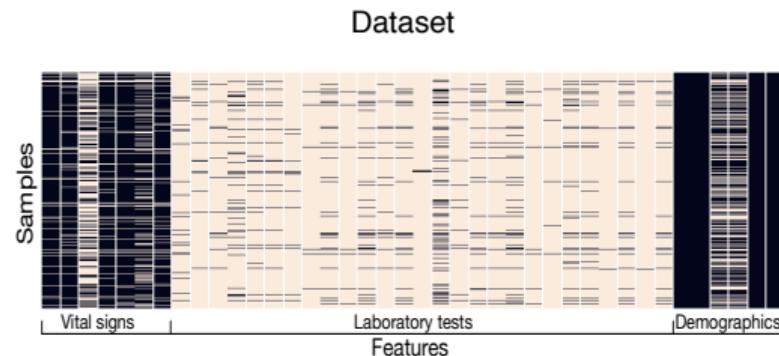
Novel Imputing Method and Deep Learning Techniques for Early Prediction of Sepsis in ICU



- Challenge: predict sepsis 6 hours before the clinical prediction of sepsis
- 3 ICUs
- +2000 patients

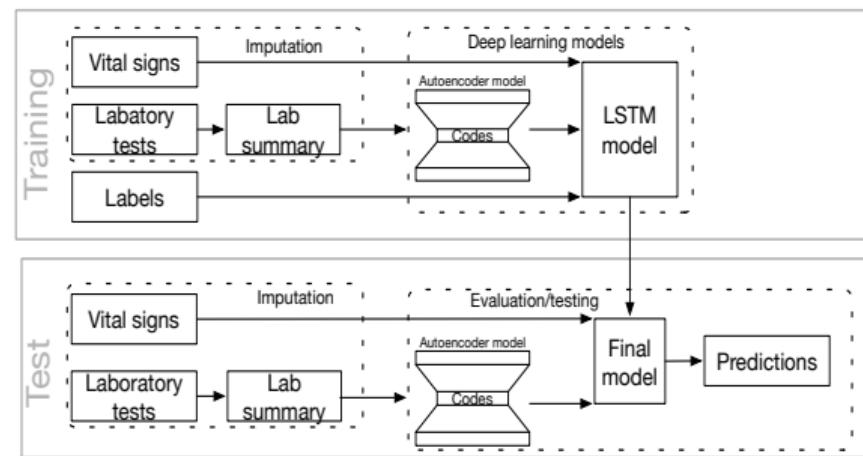
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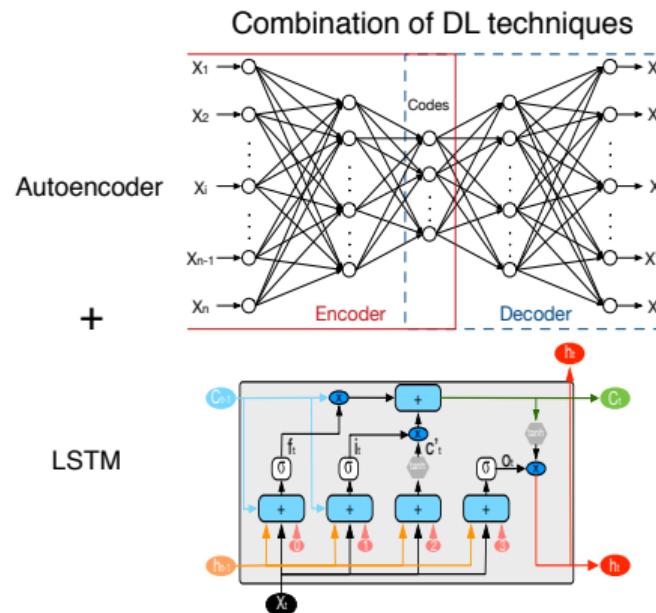
Proposed method: Combine DL techniques for imputation and prediction





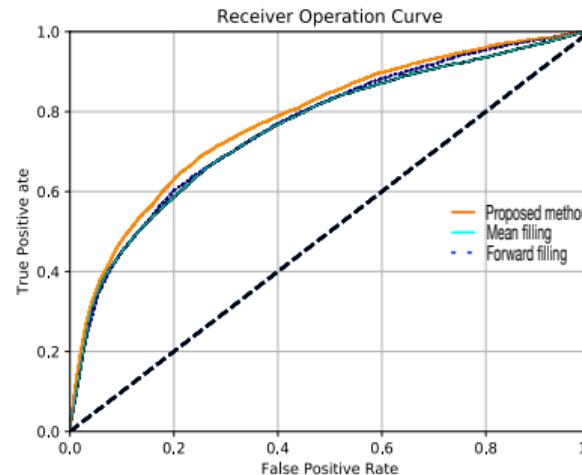
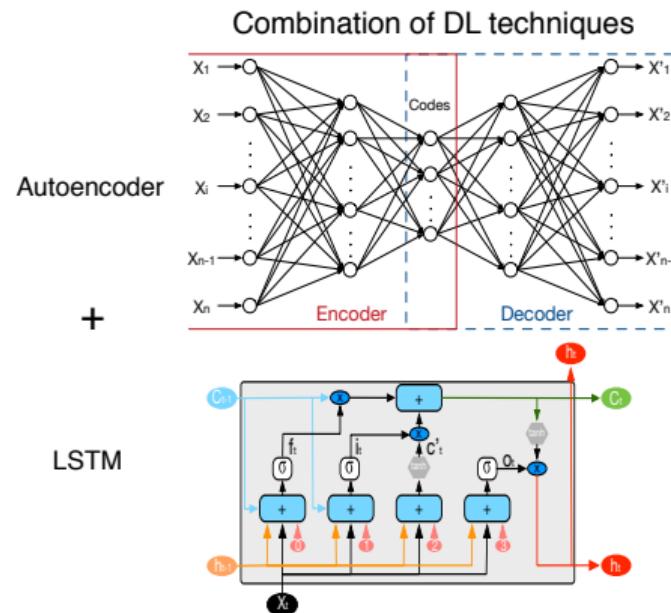
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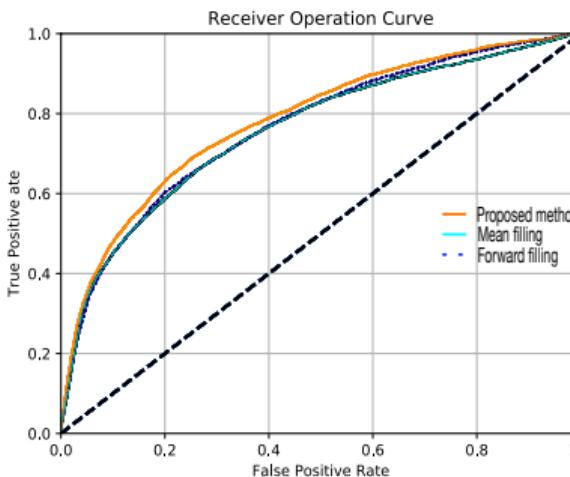
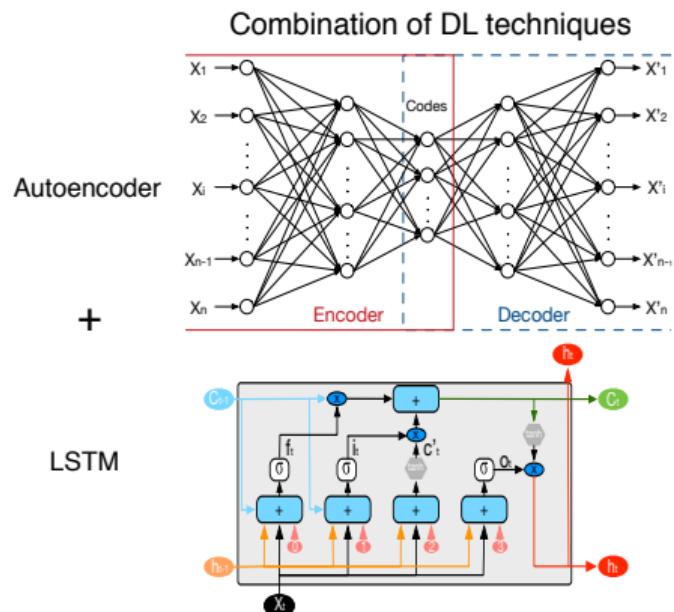


Results

- 38/78 in challenge
- 27% dimensionality reduction
- 10% better in Utility

Application to EHR + Physiological Signals

Novel Imputing Method and Deep Learning Techniques for Early Prediction of Sepsis in ICU



Results

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Sumary

The Evolution of Machines

Data Analytics in Health

Application to Physiological Signals: Arousal detection

Application to EHR

Application to EHR + Physiological Signals

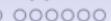
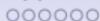
Practical Information on the Course



Goals

Applied Machine Learning

- Understand the intuition behind modern machine learning methods as well as a bit more formal understanding of how, why, and when they work.
- Apply the machine learning techniques over data to solve practical problems.



Contents

Applied Machine Learning

1. Fundamentals of ML item Data Pre-processing
2. Learning from data
3. Supervised learning techniques
4. Non-supervised learning techniques
5. Artificial Neural Networks

Bibliography

Applied Machine Learning

1. *Hands on Machine Learning*, A. Geron. 2/3th edition

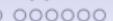
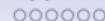
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VLE + Teams

Applied Machine Learning

The course is based on the Virtual Classroom.

- Communication
- Materials
- Deliveries
- Marks



Evaluation

Applied Machine Learning

- Sitting exam: 60%
- Assignments: 30%
- Capstone project: 10%