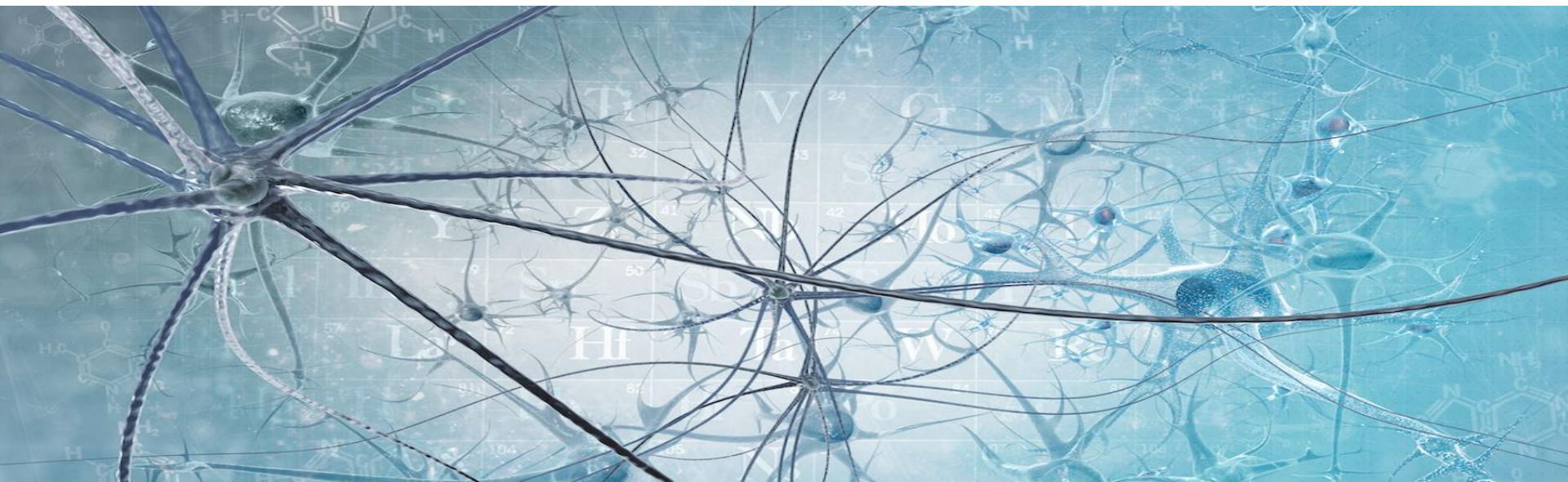


European Biotech Week  
Pavia, Sept 25<sup>th</sup> 2019

# Artificial Intelligence in Healthcare and Life Science

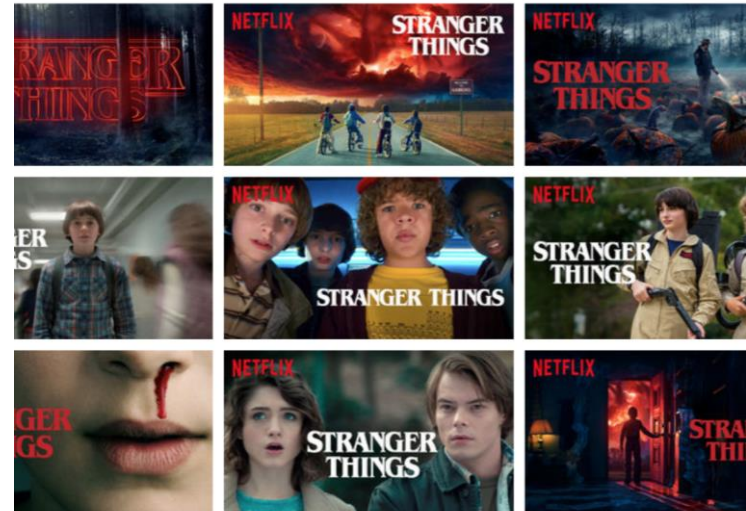
**Antonio Pelliccia** | Healthcare & Life Sciences, IBM



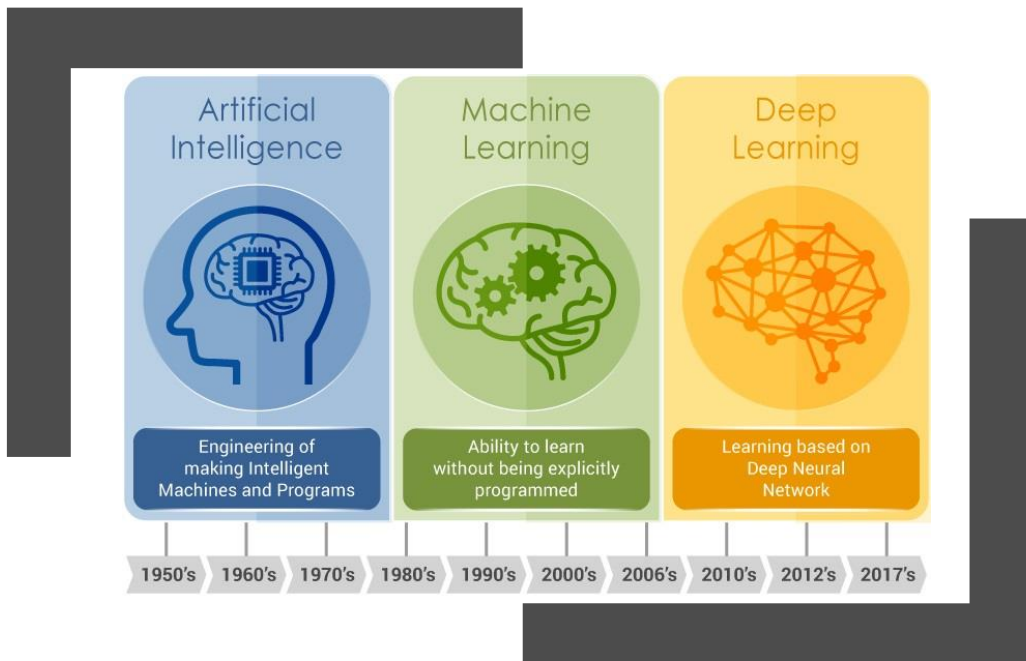
# The official birth of AI

- 1956: Darmouth Summer Research Project on Artificial Intelligence

McCarthy, Marvin Minsky, Nathaniel Rochester and Claude Shannon



# What is AI?



“**Machine learning** is a field of computer science that gives computer systems the ability to "learn" from data, **without being explicitly programmed**”

Samuel, Arthur L. (1959). "Some Studies in Machine Learning Using the Game of Checkers". *IBM Journal of Research and Development*

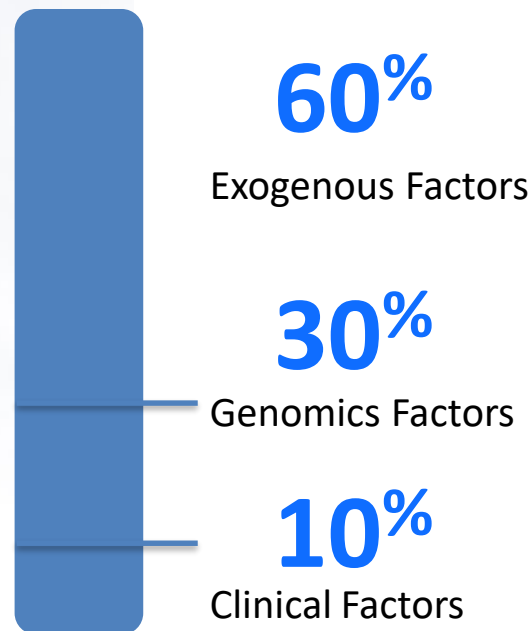


The background of the slide is a dark blue field filled with numerous thin, wavy, light blue lines that create a sense of movement and depth. Scattered throughout this field are many out-of-focus, glowing spheres in shades of yellow and white, resembling bokeh from distant lights or data points in a network. The overall effect is one of a complex, interconnected system, possibly representing neural networks or data flow.

Why is Deep  
Learning stepping  
in the limelight?

# Healthcare Data are exploding

(from 153 Exabytes in 2013 to 2.314 Exabytes in 2020 - IDC)



Source: Health Policy Brief: The Relative Contribution of Multiple Determinants to Health Outcomes," Health Affairs, August 21, 2014

If it walks/swims/quacks like a Duck... then it must be a Duck



Swims



Walks

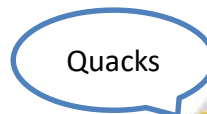


Quacks

Duck



Walks



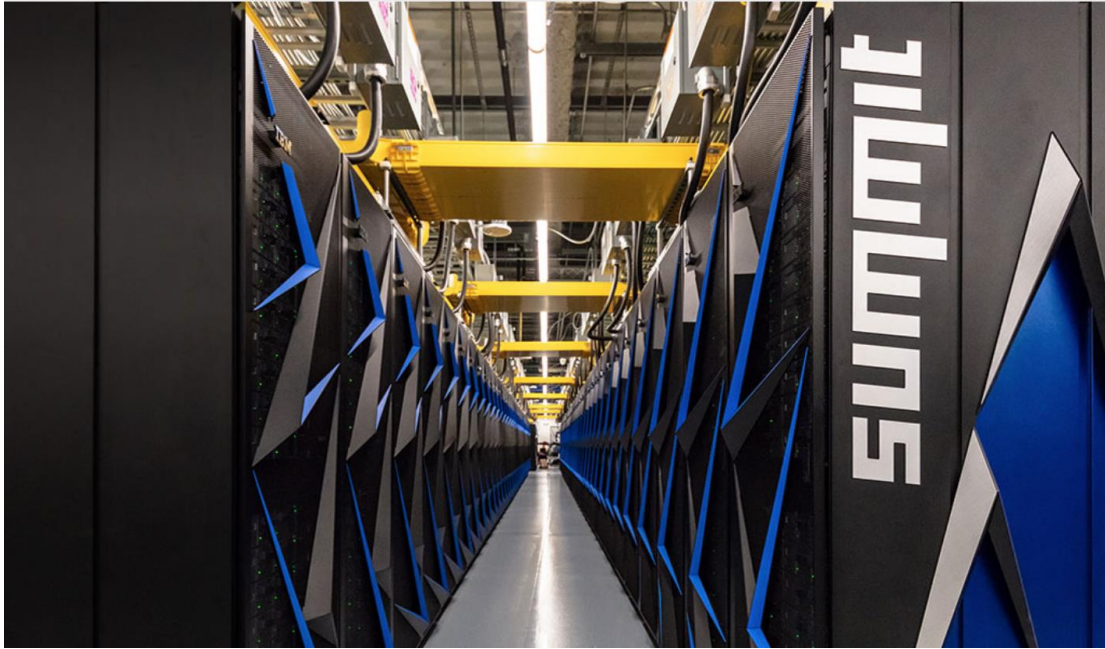
Quacks



Swims

Not a Duck





IBM builds the world's fastest  
supercomputers  
What will we do with 200 petaflops?

- Machine learning algorithms scaled on Summit will help medical researchers with a comprehensive view of the cancer population
- Using a mix of AI techniques, researchers will be able to identify patterns in the function, cooperation, and evolution of human proteins and cellular systems

# Negation disambiguation in medical text is still a challenge

A 12-year old girl with known hyperagglutinability, presented to the emergency department with a 2-week history of headaches and facial weakness. Neurologic examination indicated sensorineural hearing loss on the right side with Weber's test lateralizing to the left, and the Rinne's test demonstrating bone conduction greater than air conduction on the right. Magnetic resonance imaging of the head revealed severe structural defects of the right petrous temporal bone. No indication of cerebral infarction.

Furthermore, IL-2 and IL-12 synergistically induced IRF-1, whereas IFN-alpha and IL-12 did not.

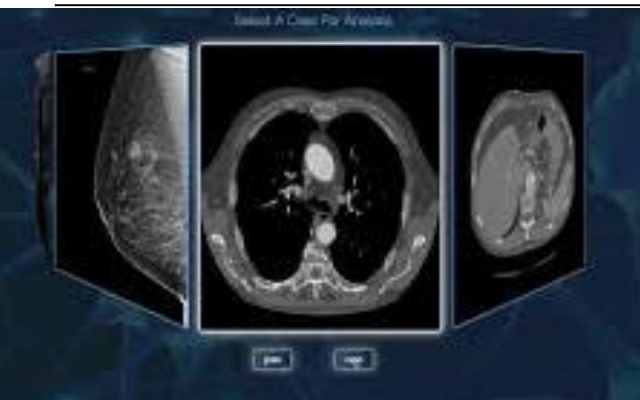
ID: E1  
TRIGGER: *synergistically induced*  
TYPE: POSITIVE REGULATION  
THEME: *IRF-1* : protein molecule  
CAUSE: *IL-2* : protein molecule  
CAUSE: *IL-12* : protein molecule  
Polarity: *Positive*

ID: E2  
TRIGGER: *synergistically induced*  
TYPE: POSITIVE REGULATION  
THEME: *IRF-1* : protein molecule  
CAUSE: *IFN-alpha* : protein molecule  
CAUSE: *IL-12* : protein molecule  
Polarity: *Negative*





It is hard to train AI for Image recognition (!?)



2010 and earlier

## Narrow AI

Single Task, Single Domain  
Superhuman accuracy and speed for certain tasks

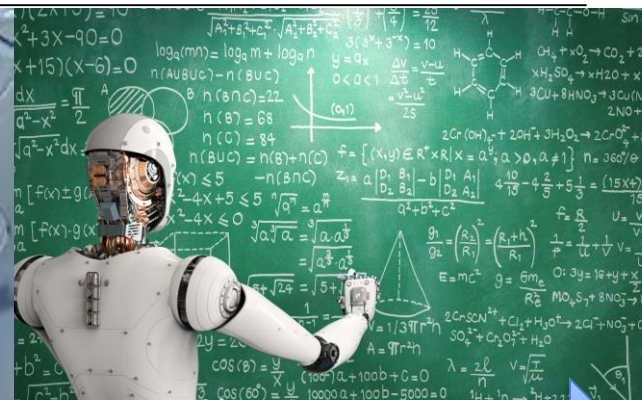
2015



2019

## Broad AI

Multi Task, Multi Domain  
Multi modal, explainable, Distributed



2050 and beyond

## General AI

Cross Domain learning and reasoning  
Broad autonomy

# Where we are nowadays?

# The evolution of AI

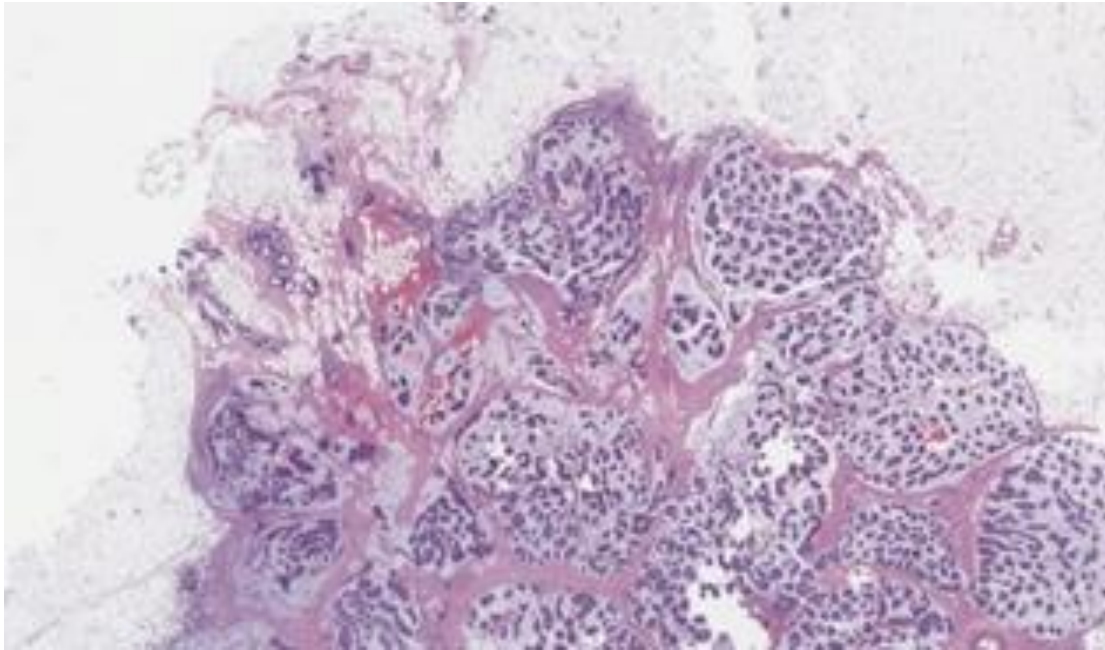


What can  
Artificial  
Intelligence  
do?



AI can understand and extract data from Natural Language:

- Literature
  - PDTA and guidelines
  - Medical data
  - Patient generated data
- 
- At Baylor College, Watson analysed 70.000 publications on p53 in few minutes and predicted proteins that turn on/off p53's activity
  - 6 potential proteins were identified for new research targets



AI can understand  
Images

Project to help anatomy  
pathologists to early  
identify precancerous  
lesions of breast tumours  
(UDH, ADH FEA, DCIS)



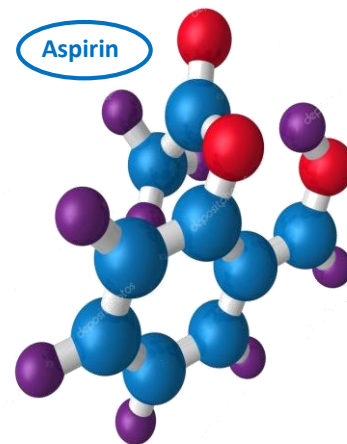
ISTITUTO NAZIONALE TUMORI  
IRCCS - Fondazione Pascale



IBM **Research** | Zurich

# A future quantum processor could simulate a drug molecule

This would require a conventional computer larger than 10 percent of the size of the earth



Type of Scaling	Time to Solve Problem				
	10 secs	2 mins	330 years	3300 years	Age of the Universe
Classical algorithm with exponential runtime	10 secs	2 mins	330 years	3300 years	Age of the Universe
Quantum algorithm with polynomial runtime	1 min	2 mins	10 mins	11 mins	~24 mins



# Use case of Artificial Intelligence for Biotech industry



AI for drug target  
identification  
and validation



AI for target based  
and phenotypic  
drug discovery



AI for dealing with  
biomedical, clinical  
and patient data



AI for  
polypharmacology  
discovery



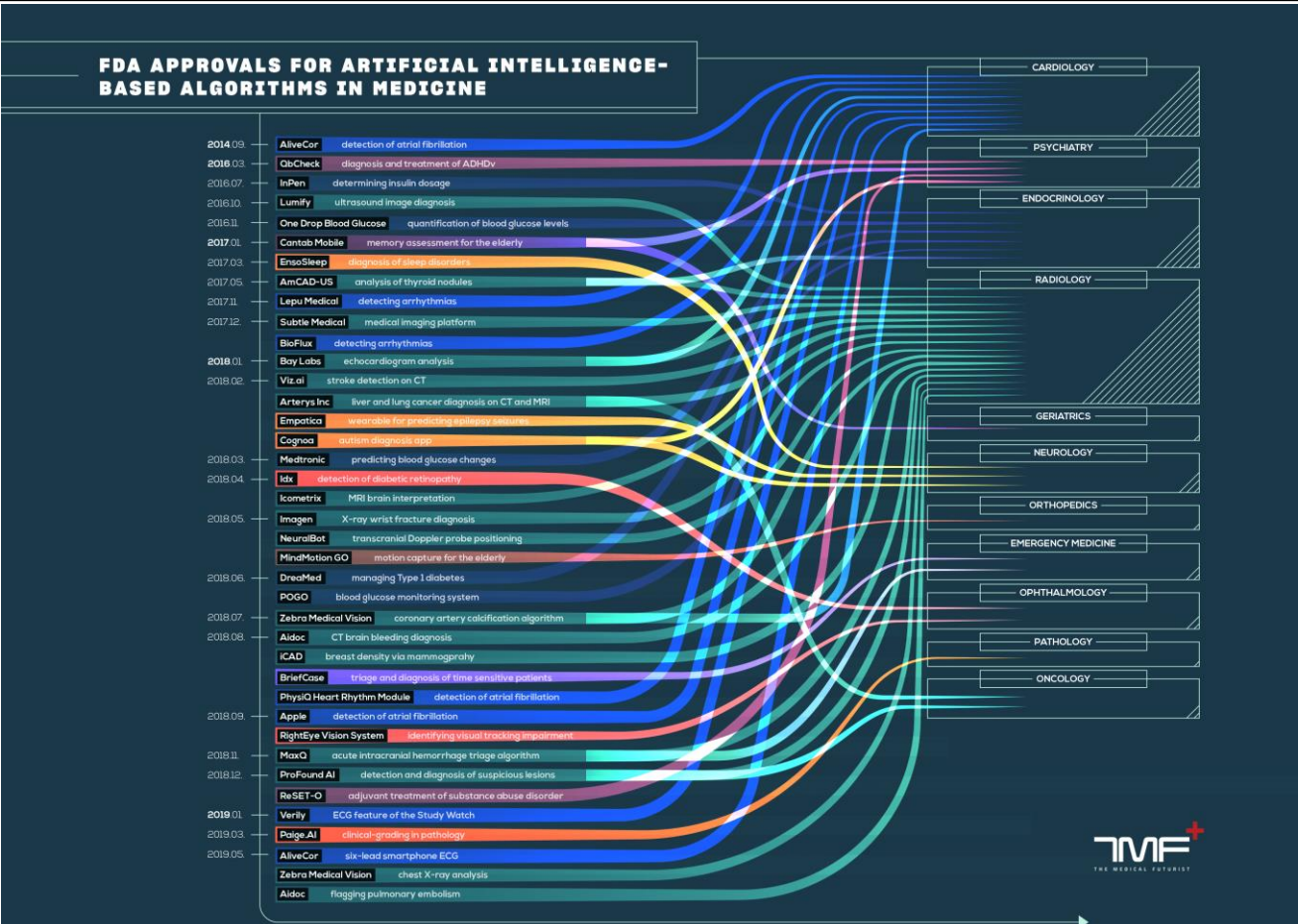
AI for drug  
repurposing  
programs



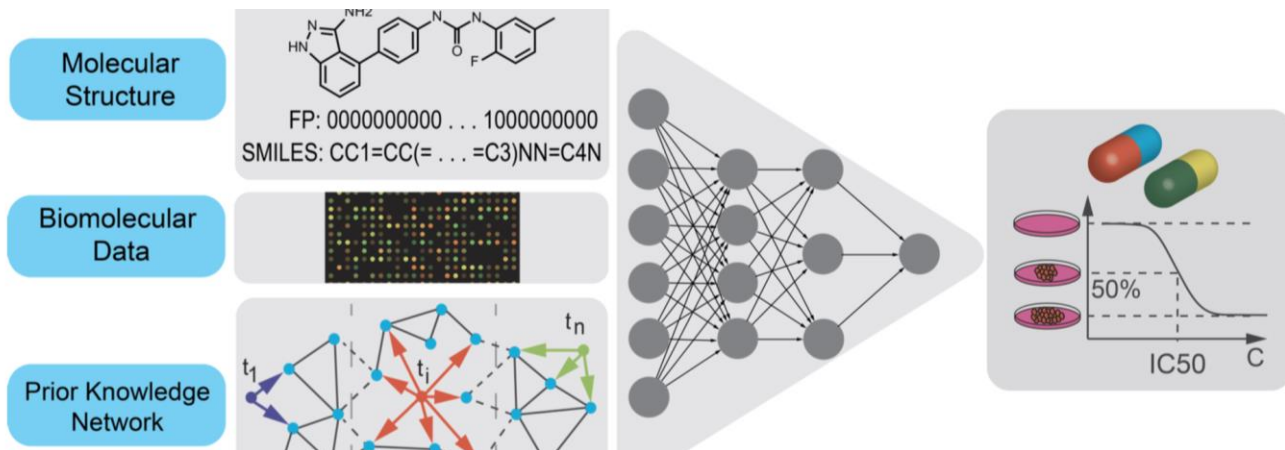
AI for  
biomarkers  
development



AI for analyzing research  
literature, publications,  
and patents



# Multi-modal prediction of IC50 drug sensitivity (PaccMann)

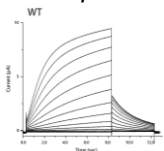


Drug	Cell line	Cancer type	Top-5 attended genes	IC50	
				Predicted	Measured
Afatinib	UMC-11	lung (NSCLC)	F13A1, MYH4, ATOH8, SEMA4A, NES	0.505	0.493
BX-912	YH-13	glioma	RNASE2, HOXA13, CBR3, FABP1, HDC	0.532	0.5
GSK319347A	EW-12	bone	CD300A, RHBDL2, NES, TFF3, SOCS1	0.597	0.7
JW-7-24-1	OVTOKO	ovary	HDC, EIF2A, RNASE2, ANGPTL6, CBR3	0.502	0.49
PI-103	MV-4-11	leukemia	TFF3, ATOH8, RBP2, ITIH3, GRIP1	0.362	0.33
TGX221	SW962	urogenital system	CBR3, RNASE2, FABP1, HDC, SH3D21	0.621	0.66
S-Trityl-L-cysteine	NCI-H187	lung (SCLC)	RHBDL2, NR1H4, MYH4, NES, APCS	0.535	0.502
Fedratinib	BL-41	lymphoma	TFF3, ATOH8, RBP2, MAPK7, ARHGEF33	0.382	0.428
Tipifarnib	RCC10RGB	kidney	EIF2A, HDC, CBR3, PIK3R5, HOXA13	0.542	0.544
Midostaurin	GAK	skin	SVOP, FABP1, HDC, F13A1, FGFR3	0.507	0.477

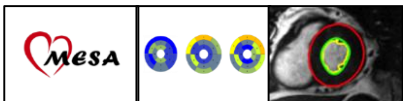


# Multiscale heart modelling for diagnosis, patient monitoring and system pharmacology

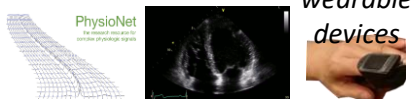
## In vitro experiments, drug effects



## Medical images (CT, MRI, US)

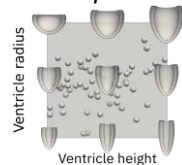


## Clinical data (ECG, BP, echo)

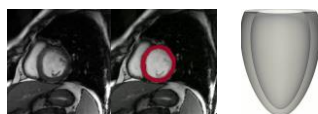


wearable  
devices

## Stochastic optimization

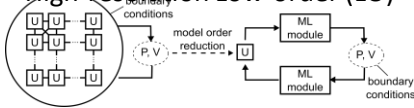


## Segmentation, parameterization

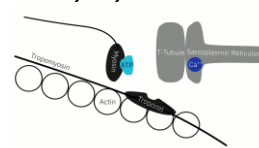


## Model-order reduction

### High-resolution Low-order (LO)



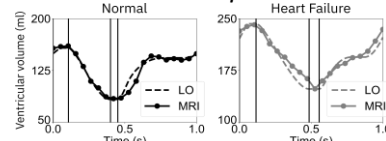
## Myocyte model



## Heart model



## Global endpoints



- Allow comprehensive exploration of cardiac drug action from cell to whole-organ and systemic levels
- Allow synthesizing virtual patient populations to better design and analyze early stage clinical trials (accounting for confounding factors)

# Using Machine Learning to Develop Blood Test For Key Alzheimer's Biomarker

It is recognized that Alzheimer's disease (AD) exists before dementia is present and that shifts in amyloid beta occur long before clinical symptoms can be detected.

IBM Research developed a blood-based signature that can provide a cheap and minimally invasive estimation of an individual's CSF amyloid status using a machine learning approach.

This is the first study to show that a machine learning approach, using plasma protein levels, age and APOEε4 carrier status, is able to predict CSF Aβ<sub>1-42</sub> status, the earliest risk indicator for AD, with high accuracy.



Article | OPEN | Published: 11 March 2019

## A blood-based signature of cerebrospinal fluid Aβ<sub>1-42</sub> status

Benjamin Goudey, Bowen J. Fung, Christine Schieber, Noel G. Faux<sup>✉</sup>, Alzheimer's Disease Metabolomics Consortium & Alzheimer's Disease Neuroimaging Initiative

Scientific Reports 9, Article number: 4163 (2019) | Download Citation ↓

<https://www.nature.com/articles/s41598-018-37149-7>

Watson for  
Clinical Trials  
Matching  
matches patients  
to open clinical  
trials







*AI is helping Barrow Neurological Institute narrow research scope and uncover new pathways of interest for drug therapies in the fight against ALS:*

- **5 new proteins identified in months***
- **80% of top-ranked targets were proven to be linked to ALS***
- **Identifies new pathways of interest for drug therapies** that scientists may not have considered otherwise*





# Predict the best second level therapy to type-2 diabetic patients using RWE



## Estimating the effects of second-line therapy for type 2 diabetes mellitus: retrospective cohort study

Gottlieb A, Yanover C, Cahan A, Goldschmidt Y. *BMJ Open Diab Res Care* 2017;5:e000435.



### Causal inference methods applied to observational data aligned with current evidence

Predicted reduction in glycosylated hemoglobin (HbA1c) levels for sulfonylureas, the most commonly prescribed second-line drugs, was smaller than for other tested classes (TZDs, GLP-1 agonists, DPP-4 inhibitors)

Predicted significant reduction of body mass index with DPP-4 inhibitors compared to sulfonylureas and TZDs

\*TZDs = thiazolidinediones, GLP-1 = glucagon-like peptide-1, DPP-4 = dipeptidyl peptidase 4

*"EHR data can support causal inference and allow replication of clinical trial results. The advantages of this approach in terms of the labor and costs required to expand evidence-based medicine are clear."*

# Artificial Intelligence supports Oncologic Genomics

- Watson for Genomics analyses tumor sample sequencing and in 2 minutes returns a report with all actionable mutations that can be addressed by an existent therapy or a clinical trial with all the relevant literature references
- In a retrospective study of about 1000 cases, Watson for Genomics has achieved a concordance rate of 99%, highlighting new therapeutic options coming from recent studies for about 33% of the sample

# With Watson you can transform personal healthcare

415 million adults with diabetes in the world – a number that could grow to 642 million by 2040

Cost of diabetes and prediabetes in US at \$322 billion

Sugar.IQ with Watson app features real-time, continuous glucose monitoring (CGM) to predict hypoglycemia 2 to 3 hours in advance, with an accuracy of 85-89%

With Watson, insights drive positive behavior changes among app users to control blood sugar levels



Medtronic





# aimac

informa per aiutare  
a vivere con il cancro

IBM has developed a virtual assistant that helps the citizens and caregivers to ask questions about oncologic patient rights



IBM collaborated with Cooperativa Sole to develop new technologies for innovative care models

**sole.**  
persone per le persone



The best technology... is the invisible one



A healthcare worker in blue scrubs is using a stethoscope to check the back of an elderly woman sitting on a bed. The woman has grey hair and is wearing a patterned top. The scene is set in a home with a couch and a lamp visible in the background. The text "Thanks for your attention" is overlaid in white.

Thanks for your  
attention