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2019

## Artificial Intelligence in Cardiology and Cardiac Surgery

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# What is “Intelligence”?

- “. . . the ability of a system to act appropriately in an uncertain environment, where appropriate action is that which increases the probability of success, and success is the achievement of behavioral subgoals that support the system’s ultimate goal.” J. S. Albus [1]
- “Any system . . . that generates adaptive behaviour to meet goals in a range of environments can be said to be intelligent.” D. Fogel [10]
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- “Achieving complex goals in complex environments” B. Goertzel [12]
- “Intelligent systems are expected to work, and work well, in many different environments. Their property of intelligence allows them to maximize the probability of success even if full knowledge of the situation is not available. Functioning of intelligent systems cannot be considered separately from the environment and the concrete situation including the goal.” R. R. Gudwin [15]
- “[Performance intelligence is] the successful (i.e., goal-achieving) performance of the system in a complicated environment.” J. A. Horst [18]
- “Intelligence is the ability to use optimally limited resources – including time – to achieve goals.” R. Kurzweil [20]
- “Intelligence is the power to rapidly find an adequate solution in what appears a priori (to observers) to be an immense search space.” D. Lenat and E. Feigenbaum [23]
- “Intelligence measures an agent’s ability to achieve goals in a wide range of environments.” S. Legg and M. Hutter [21]
- “...doing well at a broad range of tasks is an empirical definition of ‘intelligence’ ” H. Masum [24]
- “Intelligence is the computational part of the ability to achieve goals in the world. Varying kinds and degrees of intelligence occur in people, many animals and some machines.” J. McCarthy [25]
- “. . . the ability to solve hard problems.” M. Minsky [26]
- “Intelligence is the ability to process information properly in a complex environment. The criteria of properness are not predefined and hence not available beforehand. They are acquired as a result of the information processing.” H. Nakashima [27]
- “. . . in any real situation behavior appropriate to the ends of the system and adaptive to the demands of the environment can occur, within some limits of speed and complexity.” A. Newell and H. A. Simon [29]
- “[An intelligent agent does what] is appropriate for its circumstances and its goal, it is flexible to changing environments and changing goals, it learns from experience, and it makes appropriate choices given perceptual limitations and finite computation.” D. Poole [31]
- “Intelligence means getting better over time.” Schank [32]
- “Intelligence is the ability for an information processing system to adapt to its environment with insufficient knowledge and resources.” P. Wang [39]
- “. . . the mental ability to sustain successful life.” K. Warwick quoted in [4] 7
- 18. “. . . the essential, domain-independent skills necessary for acquiring a wide range of domain-specific knowledge – the ability to learn anything. Achieving this with ‘artificial general intelligence’ (AGI) requires a highly adaptive, general-purpose system that can autonomously acquire an extremely wide range of specific knowledge and skills and can improve its own cognitive ability through self-directed learning.” P. Voss [38]

- “The ability to use memory, knowledge, experience, understanding, reasoning, imagination and judgement in order to solve problems and adapt to new situations.” AllWords Dictionary, 2006
- “The capacity to acquire and apply knowledge.” The American Heritage Dictionary, fourth edition, 2000
- “Individuals differ from one another in their ability to understand complex ideas, to adapt effectively to the environment, to learn from experience, to engage in various forms of reasoning, to overcome obstacles by taking thought.” American Psychological Association [28]
- “The ability to learn, understand and make judgments or have opinions that are based on reason” Cambridge Advance Learner’s Dictionary, 2006
- “Intelligence is a very general mental capability that, among other things, involves the ability to reason, plan, solve problems, think abstractly, comprehend complex ideas, learn quickly and learn from experience.” Common statement with 52 expert signatories [13]
- “The ability to learn facts and skills and apply them, especially when this ability is highly developed.” Encarta World English Dictionary, 2006
- “...ability to adapt effectively to the environment, either by making a change in oneself or by changing the environment or finding a new one . . . intelligence is not a single mental process, but rather a combination of many mental processes directed toward effective adaptation to the environment.” Encyclopedia Britannica, 2006
- “the general mental ability involved in calculating, reasoning, perceiving relationships and analogies, learning quickly, storing and retrieving information, using language fluently, classifying, generalizing, and adjusting to new situations.” Columbia Encyclopedia, sixth edition, 2006
- “Capacity for learning, reasoning, understanding, and similar forms of mental activity; aptitude in grasping truths, relationships, facts, meanings, etc.” Random House Unabridged Dictionary, 2006
- 2
- “The ability to learn, understand, and think about things.” Longman Dictionary or Contemporary English, 2006
- “: the ability to learn or understand or to deal with new or trying situations : . . . the skilled use of reason (2) : the ability to apply knowledge to manipulate one’s environment or to think abstractly as measured by objective criteria (as tests)” Merriam-Webster Online Dictionary, 2006
- “The ability to acquire and apply knowledge and skills.” Compact Oxford English Dictionary, 2006
- “. . . the ability to adapt to the environment.” World Book Encyclopedia, 2006
- “Intelligence is a property of mind that encompasses many related mental abilities, such as the capacities to reason, plan, solve problems, think abstractly, comprehend ideas and language, and learn.” Wikipedia, 4 October, 2006
- “Capacity of mind, especially to understand principles, truths, facts or meanings, acquire knowledge, and apply it to practise; the ability to learn and comprehend.” Wiktionary, 4 October, 2006
- “The ability to learn and understand or to deal with problems.” Word Central Student Dictionary, 2006
- “The ability to comprehend; to understand and profit from experience.” Wordnet 2.1, 2006
- “The capacity to learn, reason, and understand.” Wordsmyth Dictionary, 2006
- 3
- 3 Psychologist definitions
- This section contains definitions from psychologists. In some cases we have not yet managed to locate the exact reference and would appreciate any help in doing so.
- “Intelligence is not a single, unitary ability, but rather a composite of several functions. The term denotes that combination of abilities required for survival and advancement within a particular culture.” A. Anastasi [2]
- “...that facet of mind underlying our capacity to think, to solve novel problems, to reason and to have knowledge of the world.” M. Anderson [3]
- “It seems to us that in intelligence there is a fundamental faculty, the alteration or the lack of which, is of the utmost importance for practical life. This faculty is judgement, otherwise called good sense, practical sense, initiative, the faculty of adapting oneself to circumstances.” A. Binet [5]
- 3
- “We shall use the term ‘intelligence’ to mean the ability of an organism to solve new problems . . .” W. V. Bingham [6]
- “Intelligence is what is measured by intelligence tests.” E. Boring [7]
- “. . . a quality that is intellectual and not emotional or moral: in measuring it we try to rule out the effects of the child’s zeal, interest, industry, and the like. Secondly, it denotes a general capacity, a capacity that enters into everything the child says or does or thinks; any want of ‘intelligence’ will therefore be revealed to some degree in almost all that he attempts,” C. L. Burt [8]
- “A person possesses intelligence insofar as he has learned, or can learn, to adjust himself to his environment.” S. S. Colvin quoted in [35]
- “. . . the ability to plan and structure one’s behavior with an end in view.” J. P. Das
- “The capacity to learn or to profit by experience.” W. F. Dearborn quoted in [35]
- “. . . in its lowest terms intelligence is present where the individual animal, or human being, is aware, however dimly, of the relevance of his behaviour to an objective. Many definitions of what is indefinable have been attempted by psychologists, of which the least unsatisfactory are 1. the capacity to meet novel situations, or to learn to do so, by new adaptive responses and 2. the ability to perform tests or tasks, involving the grasping of relationships, the degree of intelligence being proportional to the complexity, or the abstractness, or both, of the relationship.” J. Drever [9]
- “Intelligence A: the biological substrate of mental ability, the brain’s neuroanatomy and physiology; Intelligence B: the manifestation of intelligence A, and everything that influences its expression in real life behavior; Intelligence C: the level of performance on psychometric tests of cognitive ability.” H. J. Eysenck.
- “Sensory capacity, capacity for perceptual recognition, quickness, range or flexibility or association, facility and imagination, span of attention, quickness or alertness in response.” F. N. Freeman quoted in [35]
- “. . . adjustment or adaptation of the individual to his total environment, or limited aspects thereof ...the capacity to reorganize one’s behavior patterns so as to act more effectively and more appropriately in novel situations . . . the ability to learn . . . the extent to which a person is educable . . . the ability to carry on abstract thinking . . . the effective use of concepts and symbols in dealing with a problem to be solved . . .” W. Freeman
- “An intelligence is the ability to solve problems, or to create products, that are valued within one or more cultural settings.” H. Gardner [11]
- 4
- “...performing an operation on a specific type of content to produce a particular product.” J. P. Guilford
- “Sensation, perception, association, memory, imagination, discrimination, judgement and reasoning.” N. E. Haggerty quoted in [35]
- “The capacity for knowledge, and knowledge possessed.” V. A. C. Henmon [16]
- “. . . cognitive ability,” R. J. Herrnstein and C. Murray [17]
- “. . . the resultant of the process of acquiring, storing in memory, retrieving, combining, comparing, and using in new contexts information and conceptual skills.” Humphreys
- “Intelligence is the ability to learn, exercise judgment, and be imaginative.” J. Huarte
- “Intelligence is a general factor that runs through all types of performance.” A. Jensen
- “Intelligence is assimilation to the extent that it incorporates all the given data of experience within its framework . . . There can be no doubt either, that mental life is also accommodation to the environment. Assimilation can never be pure because by incorporating new elements into its earlier schemata the intelligence constantly modifies the latter in order to adjust them to new elements.” J. Piaget [30]
- “Ability to adapt oneself adequately to relatively new situations in life.” R. Pinter quoted in [35]
- “A biological mechanism by which the effects of a complexity of stimuli are brought together and given a somewhat unified effect in behavior.” J. Peterson quoted in [35]
- “. . . certain set of cognitive capacities that enable an individual to adapt and thrive in any given environment they find themselves in, and those cognitive capacities include things like memory and retrieval, and problem solving and so forth. There’s a cluster of cognitive abilities that lead to successful adaptation to a wide range of environments.” D. K. Simon [33]
- “Intelligence is part of the internal environment that shows through at the interface between person and external environment as a function of cognitive task demands.” R. E. Snow quoted in [34]
- “. . . I prefer to refer to it as ‘successful intelligence.’ And the reason is that the emphasis is on the use of your intelligence to achieve success in your life. So I define it as your skill in achieving whatever it is you want to attain in your life within your sociocultural context — meaning that
- 5
- people have different goals for themselves, and for some it’s to get very good grades in school and to do well on tests, and for others it might be to become a very good basketball player or actress or musician.” R. J. Sternberg [36]
- “. . . the ability to undertake activities that are characterized by (1) difficulty, (2) complexity, (3) abstractness, (4) economy, (5) adaptedness to goal, (6) social value, and (7) the emergence of originals, and to maintain such activities under conditions that demand a concentration of energy and a resistance to emotional forces.” Stoddard
- “The ability to carry on abstract thinking.” L. M. Terman quoted in [35]
- “Intelligence, considered as a mental trait, is the capacity to make impulses focal at their early, unfinished stage of formation. Intelligence is therefore the capacity for abstraction, which is an inhibitory process.” L. L. Thurstone [37]
- “The capacity to inhibit an instinctive adjustment, the capacity to redefine the inhibited instinctive adjustment in the light of imaginatively experienced trial and error, and the capacity to realise the modified instinctive adjustment in overt behavior to the advantage of the individual as a social animal.” L. L. Thurstone quoted in [35]
- “A global concept that involves an individual’s ability to act purposefully, think rationally, and deal effectively with the environment.” D. Wechsler [40]
- “The capacity to acquire capacity.” H. Woodrow quoted in [35]
- “. . . the term intelligence designates a complexly interrelated assemblage of functions, no one of which is completely or accurately known in man...” R. M. Yerkes and A. W. Yerkes [41]
- “. . . that faculty of mind by which order is perceived in a situation previously considered disordered.” R. W. Young quoted in [20]

# Recurring concepts

- 1. The ability to acquire and apply knowledge and skills
- 2. The ability to learn or understand or to deal with new or trying situations

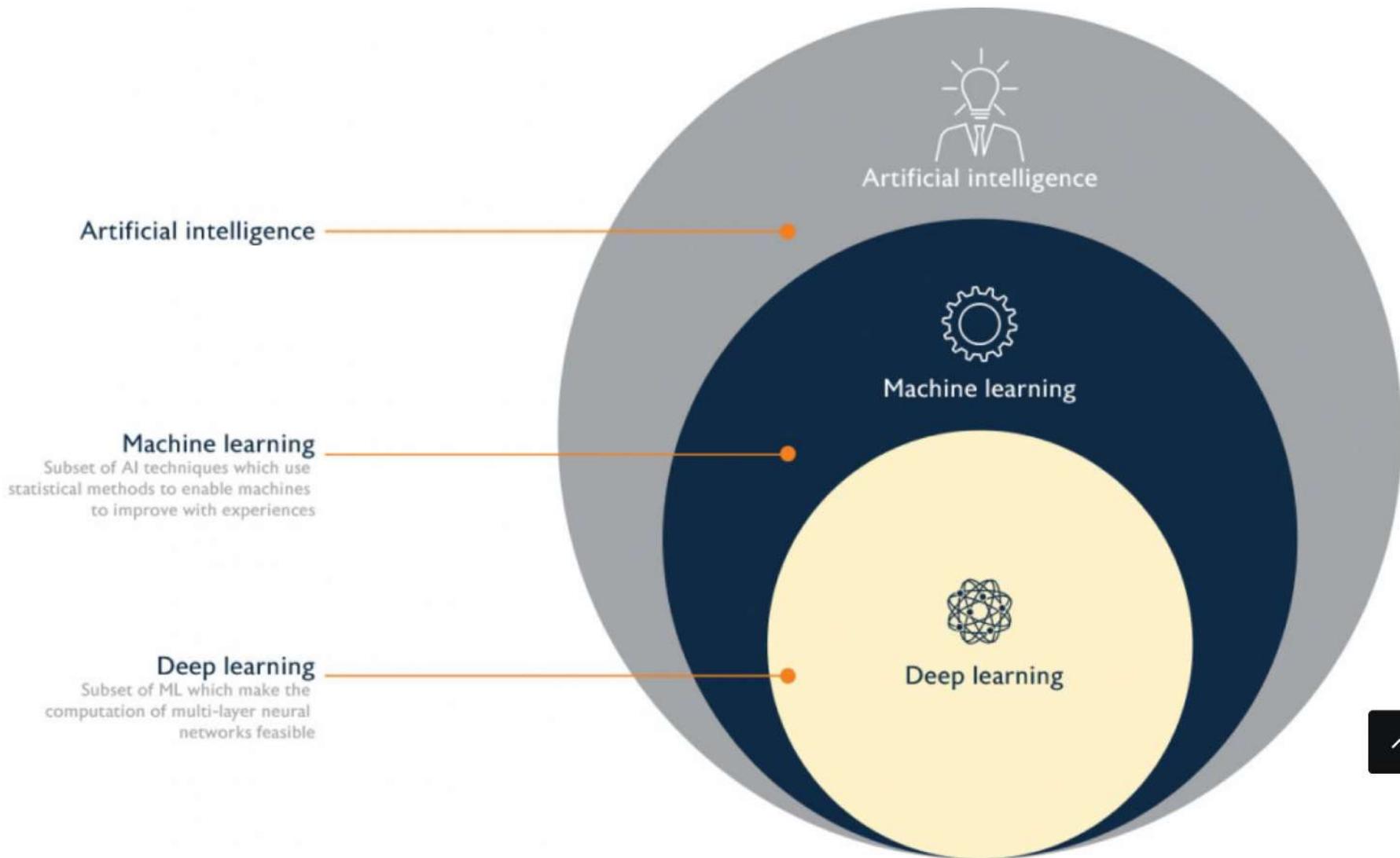
Original definition of Artificial Intelligence:

A branch of computer science that deals with the development of systems able to mimic human behavior.

TURING TEST

# AI = Ability of a computer system to:

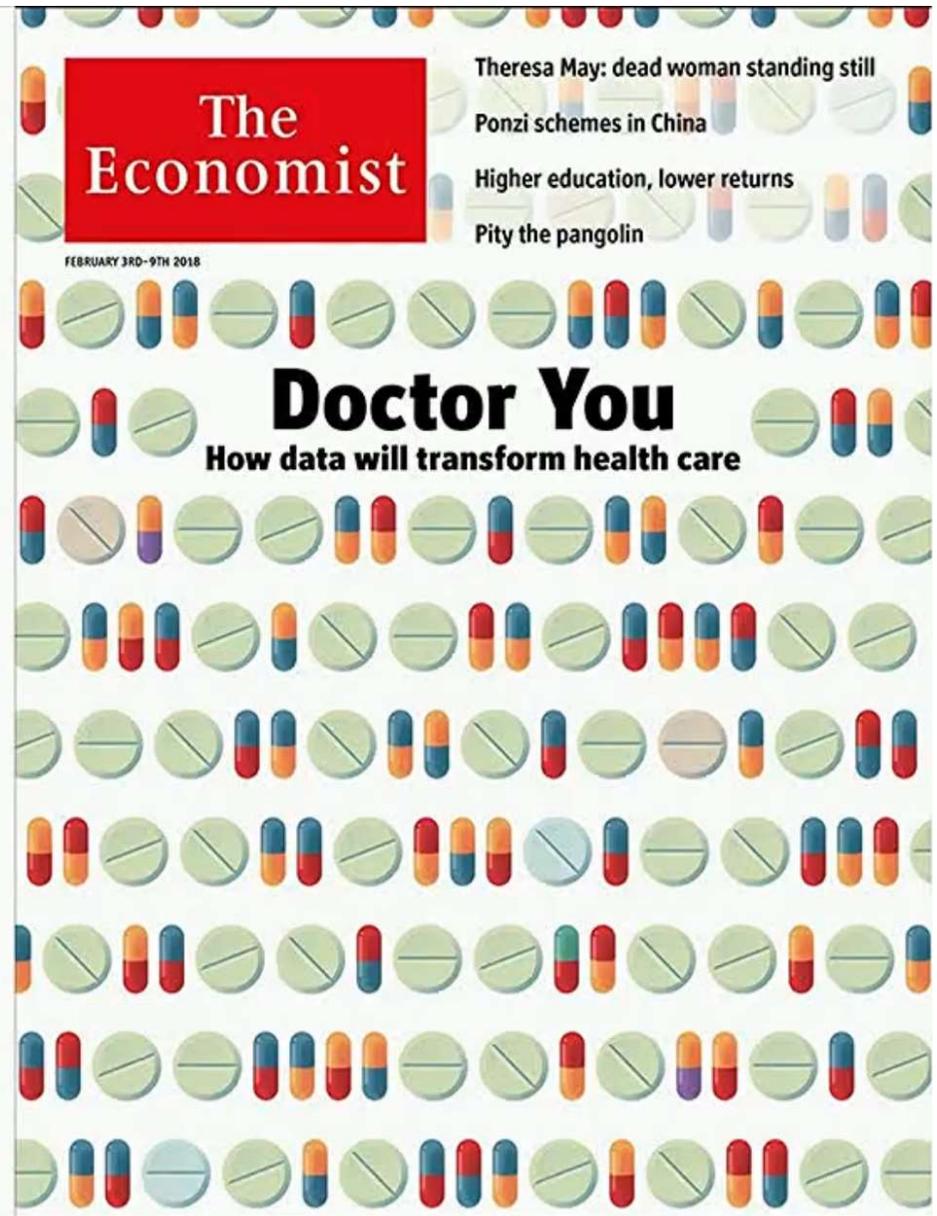
- Acquire large amounts of data (Deep Learning)
- Analyze those data with regard to the system's goal (requires training algorithms and enormous computing power / Neural Networks)
- Provide an output
- Gather feedback and use it to confirm/validate or modify the initial algorithms (Machine Learning)



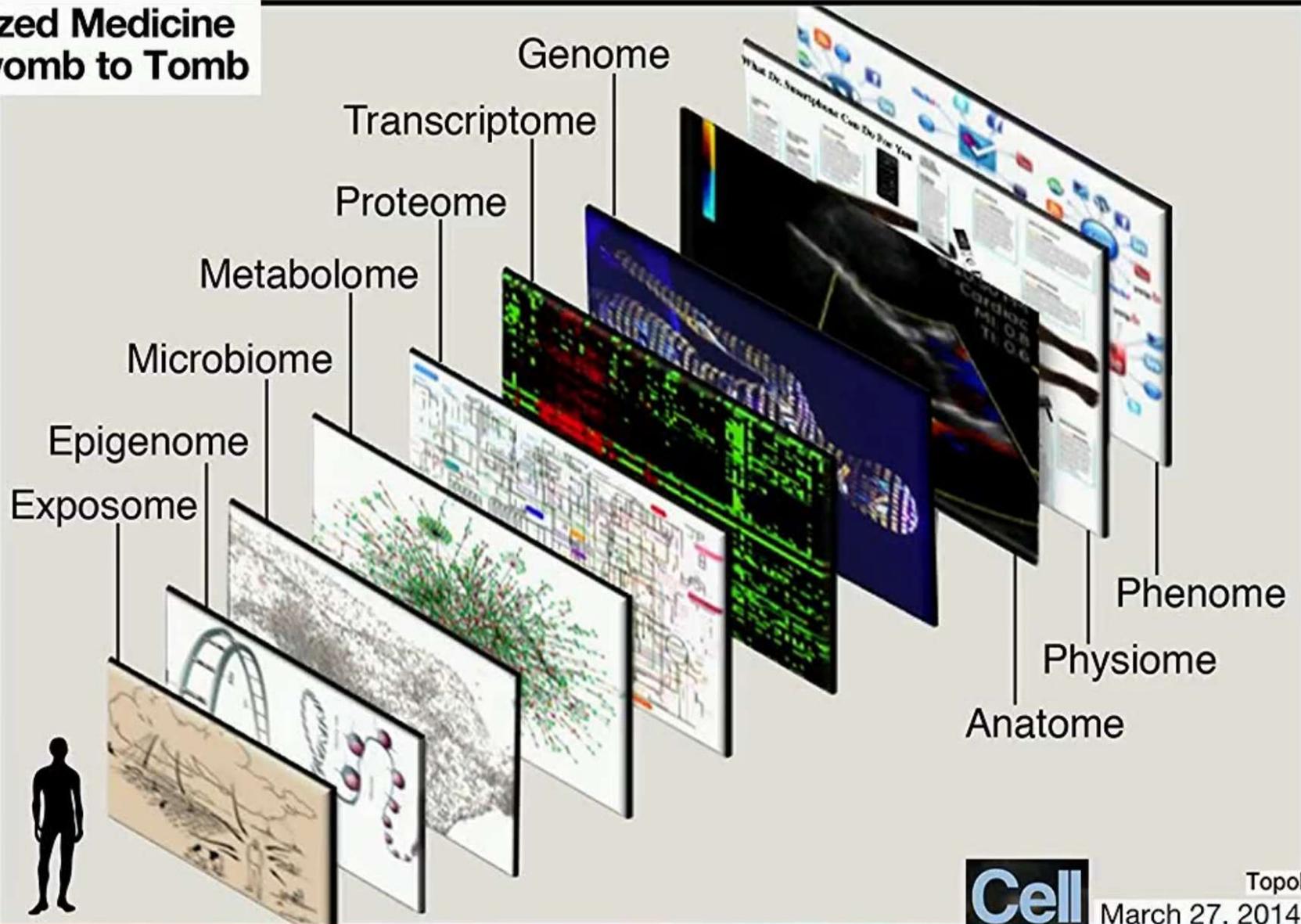


# The Importance of DATA

**Artificial Intelligence.  
Genuinely unintelligent  
without high quality data**



# Individualized Medicine from Prewomb to Tomb



Cell

Topol  
March 27, 2014

### Eye

- Glucose-sensing lens
- Digital fundoscope
- Smartphone visual-acuity tracking
- Automated refractive error
- Noninvasive intraocular pressure

### Ear

- Smart hearing aids
- Digital otoscope

### Lung

- Home spirometry
- Pulse oximetry
- Inhaler use
- Breath-based diagnostics
- Breathing sounds
- Environmental exposure

### Blood

- Continuous glucose
- Transdermal Hb
- Pathogens (genomics-based)
- PoC blood tests

### Skin

- Temperature
- Gross lesions
- Pressure sensor (wound care)
- Sweat chemistry
- Cutaneous blood flow

### Other sensors and monitors

- Pill-box and -bottle
- Posture
- Body position
- Activity
- Sleep

### Bladder and urine

- Comprehensive urinalysis
- STDs (genomic detection)
- Diaper-based sensors

### Brain and emotion

- Wireless mobile EEG
- Seizure
- Autonomic nervous activity
- Head-impact sensor
- Intracranial pressure (noninvasive)
- Stress recognition (voice, respiration)

### Heart and vascular

- Continuous BP tracking
- Handheld ECG
- Heart rhythm
- Cardiac output
- Stroke volume
- Thoracic impedance (fluid)

### Gastrointestinal

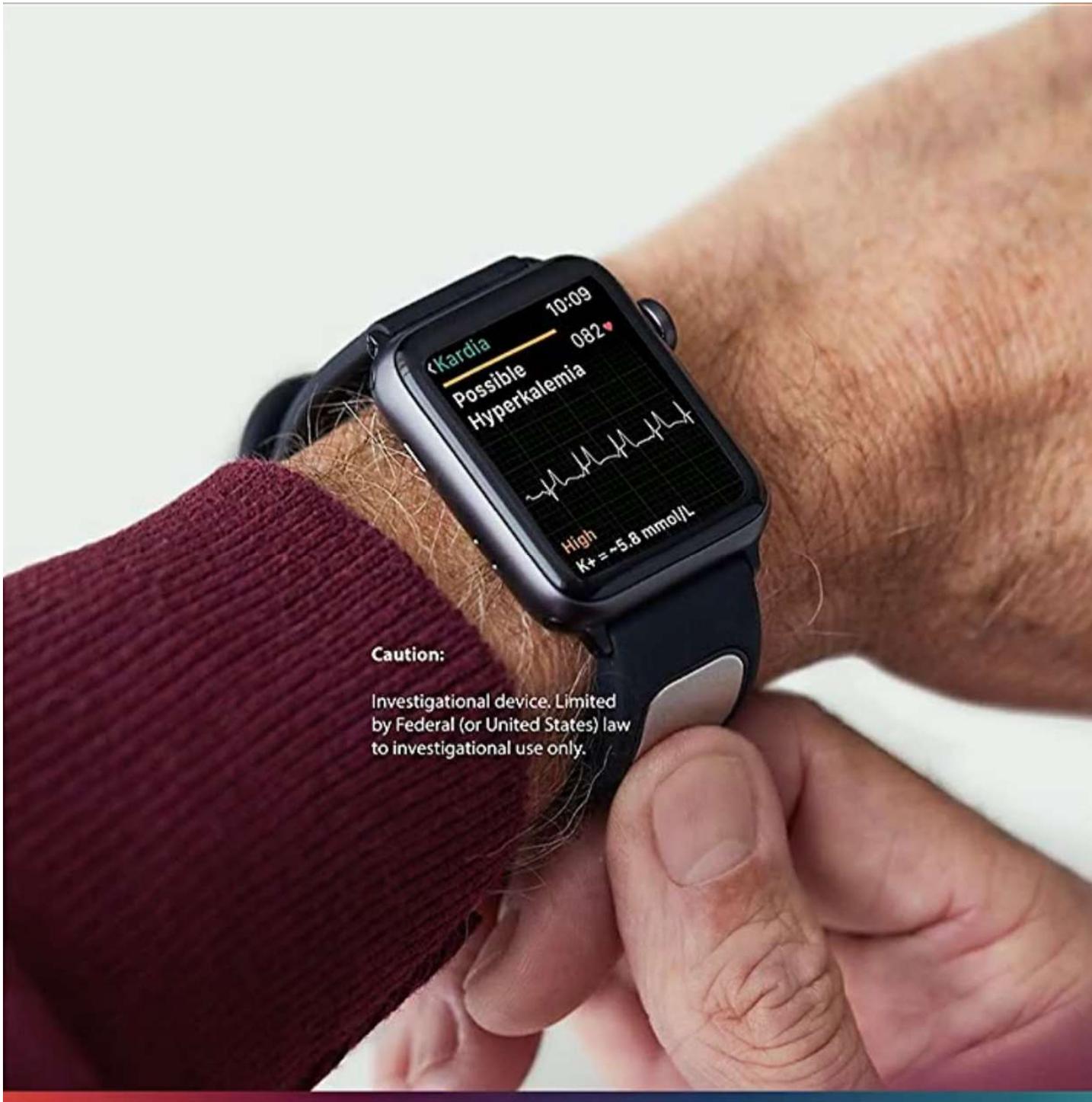
- Endoscopic imaging
- Esophageal pH
- Medication compliance
- Fecal blood or bilirubin
- Gut electrical activity
- Chewing

### Watching over one's health

- Pulse
- BP
- Temperature
- Activity
- Hydration
- Sleep stages
- Seizure
- Respiration rate
- O<sub>2</sub> saturation
- Blood CO<sub>2</sub>
- Blood glucose
- ECG (single-lead)
- Cardiac output
- Stroke volume
- Stress:
- Heart-rate variability
- Electrodermal activity

15 April 2015





**Caution:**

Investigational device. Limited by Federal (or United States) law to investigational use only.



SYS  
mmHg 127

DIA  
mmHg 84

PULSE  
/min 72

01/10 7:30 AM

# Continuous Glucose Sensors





## THE MEDICALIZED SMARTPHONE



#CARDIAC ARREST

## 'Alexa, Check My Heart', AI Tool to Detect Cardiac Arrest During Sleep

news18.com · PTI

Cardiac arrest patients are often unconscious, and have agonal breathing sounds. AI tools can help determine whether the patient needed...

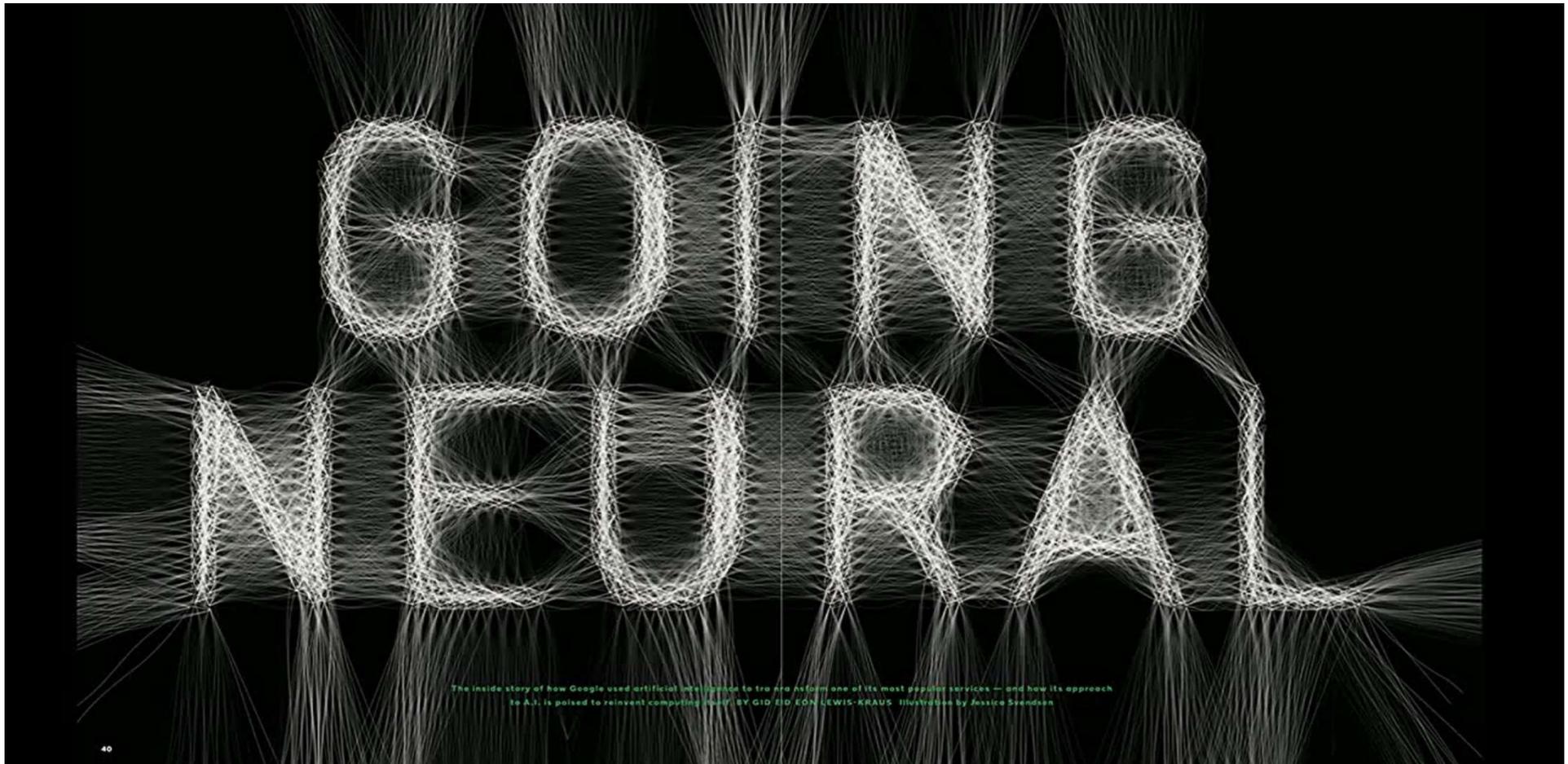
'A lot of people have smart speakers in their homes, and these devices have amazing capabilities that we can take advantage of,' said Shyam Gollakota, an associate professor at UW.

'We envision a contactless system that works by continuously and passively monitoring the bedroom for an agonal breathing event, and alerts anyone nearby to come provide CPR. And then if there's no response, the device can automatically call 911,' said Gollakota.

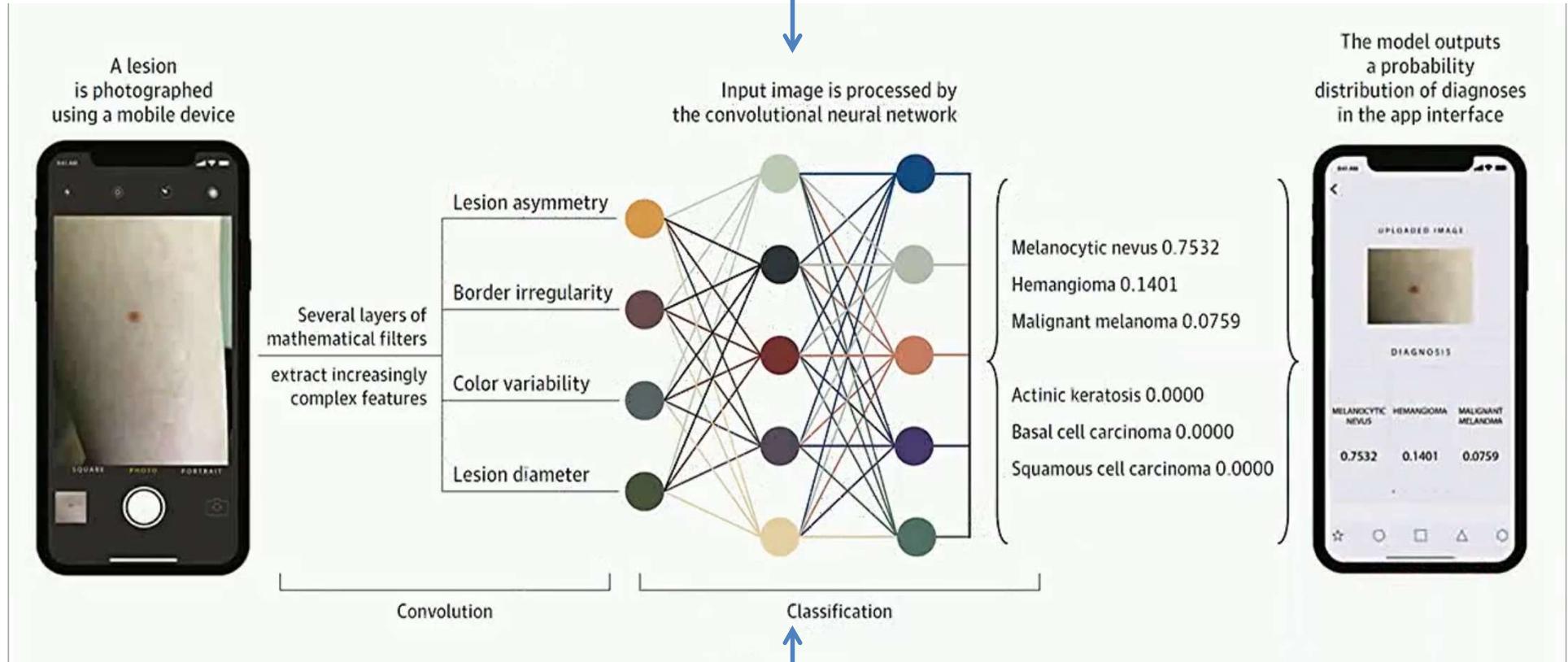
Agonal breathing is present for about 50 per cent of people who experience cardiac arrests, and patients who take agonal breaths often have a better chance of surviving.

'This kind of breathing happens when a patient

# Step 2: the analysis of data: Neural Networks



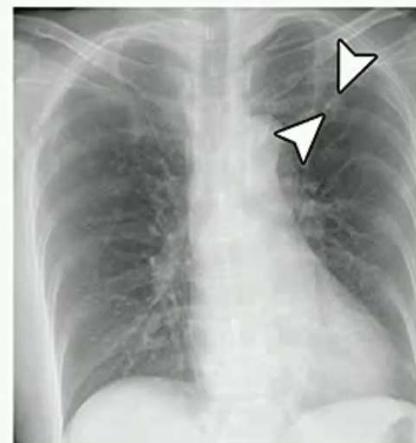
# TRAINING



FEEDBACK (Machine Learning)



34,676 patients; 43,292 Chest X-rays  
13 board-certified radiologists, 4 centers



Parameter	Radiograph Classification Performance			Nodule Detection Performance			
	AUROC	Sensitivity (%)	Specificity (%)	F1 Score (Precision, Recall)	JAFROC FOM	Sensitivity (%)*	Rate of FP Findings (%)†
Seoul National University Hospital	0.92	79.0 (94/119) [70.8, 85.4]	95 (59/62) [86.2, 98.9]	87.0 [94.9, 79.0]	0.885	69.9 (100/143) [62.0, 76.9]	0.34 [61/181]
Boramae Hospital	0.99	91.1 (112/123) [84.6, 95.1]	98 (58/59) [80.2, 100]	94.9 [99.1, 91.1]	0.924	82.0 (114/139) [74.7, 87.6]	0.30 [54/182]
National Cancer Center	0.94	71.2 (79/111) [62.1, 78.9]	100 (70/70) [93.8, 100]	83.2 [100, 71.2]	0.831	69.6 (80/115) [60.6, 77.3]	0.02 [3/181]
University of California San Francisco Medical Center	0.96	88 (78/89) [79.0, 93.1]	93 (56/60) [83.6, 97.8]	91.2 [95.1, 87.6]	0.880	75.0 (78/104) [65.8, 82.4]	0.25 [37/149]

**Conclusion:** This deep learning–based automatic detection algorithm outperformed physicians in radiograph classification and nodule detection performance for malignant pulmonary nodules on chest radiographs, and it enhanced physicians’ performances when used as a second reader.

# US Radiologists

Read 20,000 scans/yr; 50-100/day

False negative rate >25%; false positive 2%

31% are sued (almost all due to medical diagnoses)

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800 Million scans (50 Billions images) in US/yr

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30% hospitals use teleradiology, such as vRad w/ > 500 radiologists

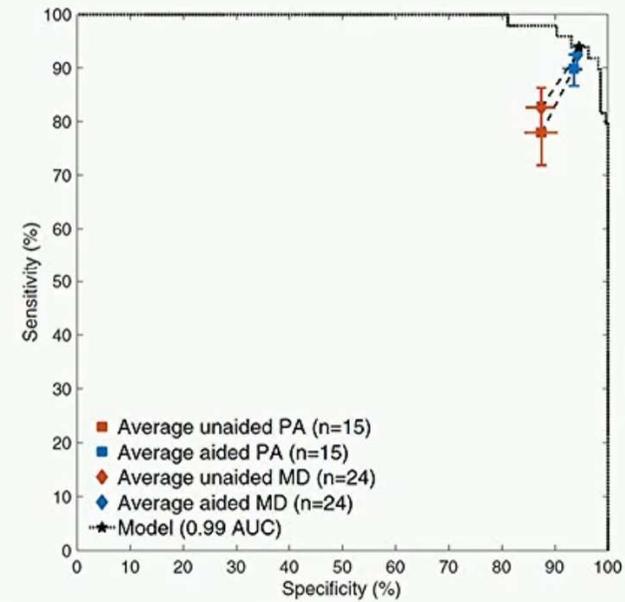
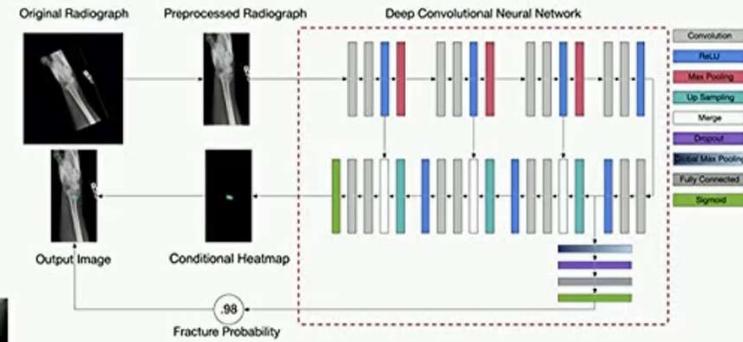
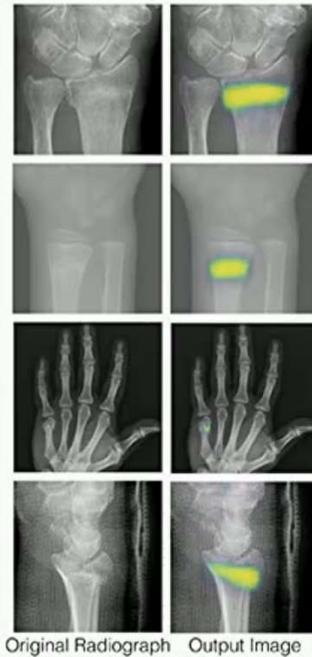


Zebra Medical, by late 2017, in 50 hospitals, > 1 million scans, \$1 dollar per scan, ~10,000 times faster



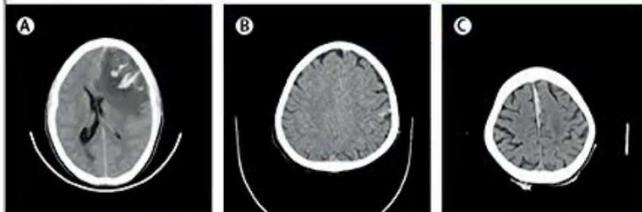
# Deep neural network improves fracture detection by clinicians

135, 409 wrist X-rays, 18 senior orthopedist  
 Tested in Emergency Medicine MDs  
 47% reduction in misinterpretation

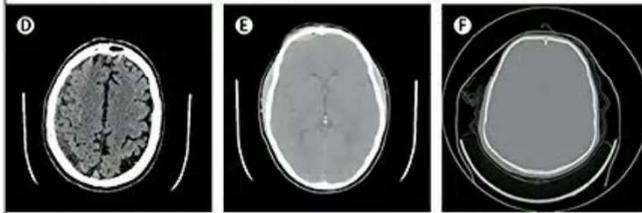


Lindsey et al,  
 22 October 2018

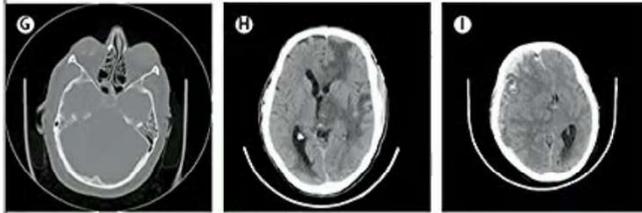
Accurate predictions: true positives



**A** Intraparenchymal haemorrhage (in left frontal region)  
**B** Subarachnoid haemorrhage (in left parietal region)  
**C** Subdural haemorrhage (along falx)

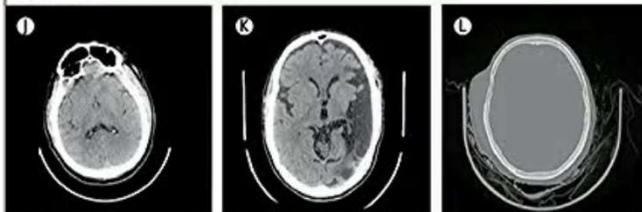


**D** Subdural haemorrhage (chronic in right parietal convexity)  
**E** Extradural haemorrhage (in right frontal convexity)  
**F** Calvarial fracture (in right parietal bone)



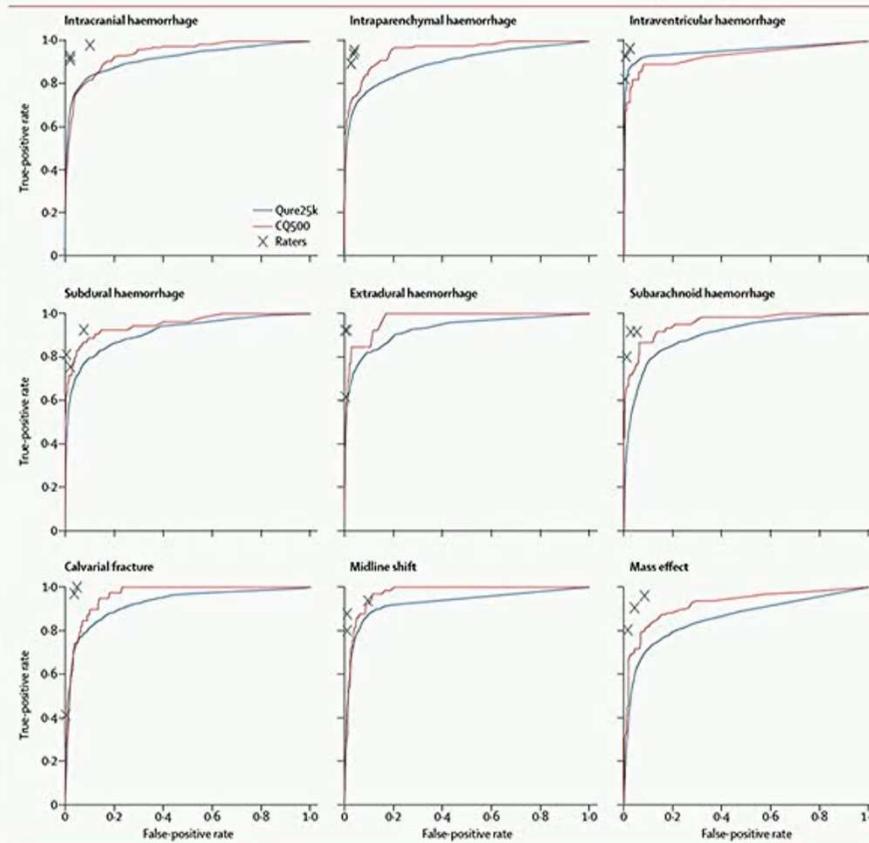
**G** Calvarial fracture (in right temporal bone)  
**H** Midline shift  
**I** Midline shift

Erroneous predictions



**J** False negative: tiny intraventricular haemorrhage  
**K** False positive: predicted as subdural haemorrhage  
**L** False negative: calvarial fracture (in right parietal bone)

# Deep learning algorithms for detection of critical findings in head CT scans: a retrospective study



313,318 Head CT scans  
 20 centers India

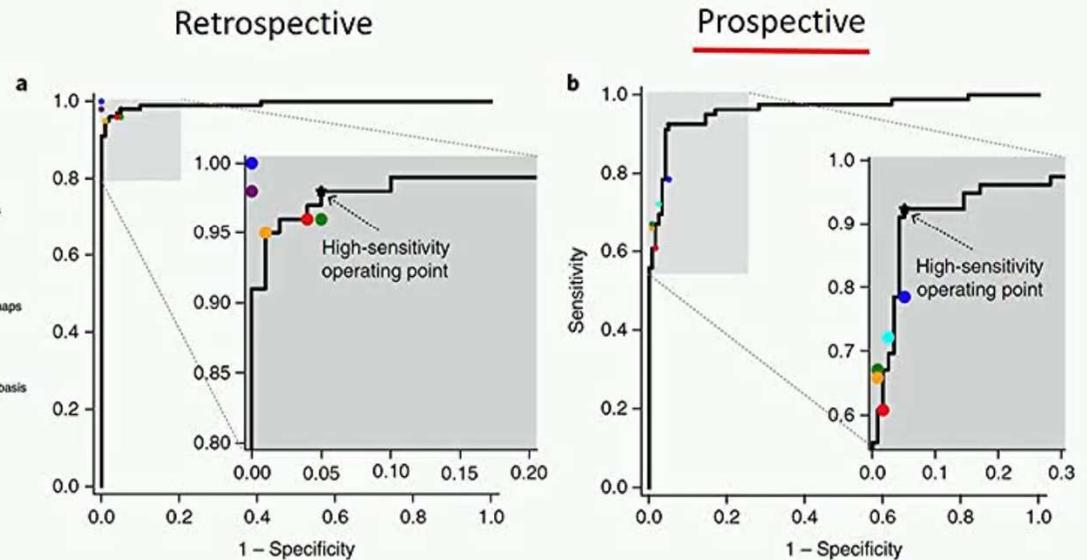
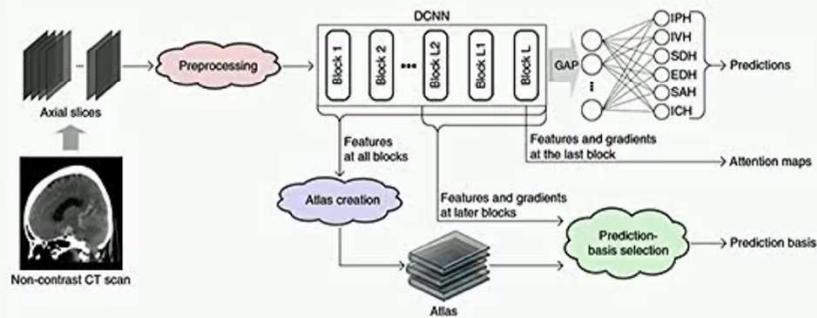
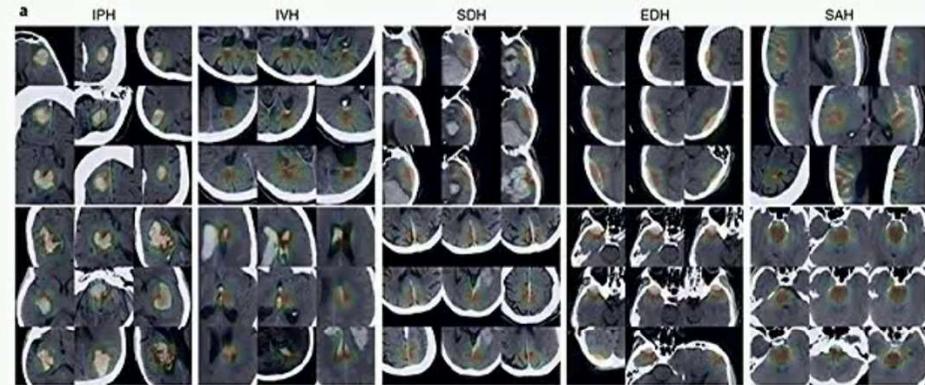
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**Interpretation** Our results show that deep learning algorithms can accurately identify head CT scan abnormalities requiring urgent attention, opening up the possibility to use these algorithms to automate the triage process.

October 11, 2018

## An explainable deep-learning algorithm for the detection of acute intracranial haemorrhage from small datasets

Hyunkwang Lee<sup>1,2,3</sup>, Sehyo Yune<sup>1,3</sup>, Mohammad Mansouri<sup>1</sup>, Myeongchan Kim<sup>1</sup>, Shahein H. Tajmir<sup>1</sup>, Claude E. Guerrier<sup>1</sup>, Sarah A. Ebert<sup>1</sup>, Stuart R. Pomerantz<sup>1</sup>, Javier M. Romero<sup>1</sup>, Shahmir Kamalian<sup>1</sup>, Ramon G. Gonzalez<sup>1</sup>, Michael H. Lev<sup>1</sup> and Synho Do<sup>1\*</sup>



# Classification and mutation prediction from non-small cell lung cancer histopathology images using deep learning

Nicolas Coudray<sup>1,2,3</sup>, Paolo Santiago Ocampo<sup>2,9</sup>, Theodore Sakellaropoulos<sup>4</sup>, Navneet Narula<sup>3</sup>, Matija Snuderl<sup>1</sup>, David Fenyo<sup>5,6</sup>, Andre L. Moreira<sup>3,7</sup>, Narges Razavian<sup>8\*</sup> and Aristotelis Tsirogas<sup>1,10\*</sup>

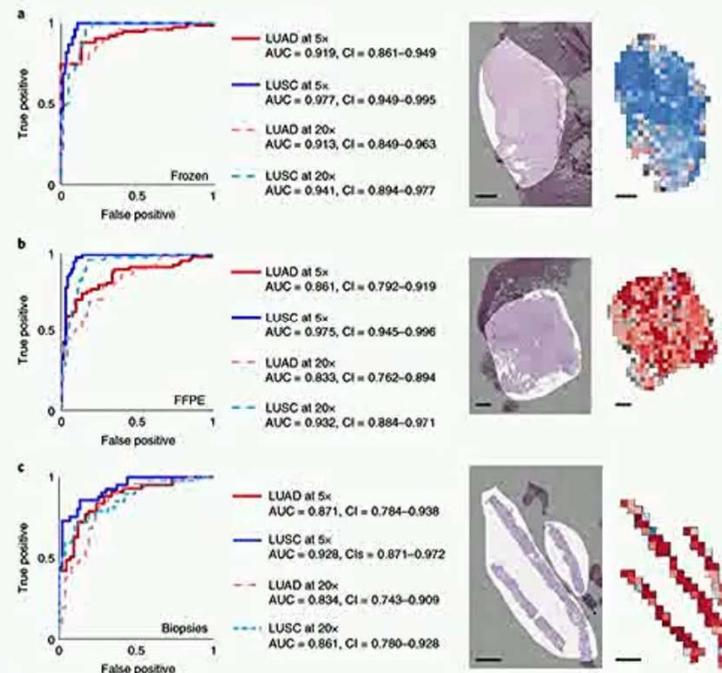
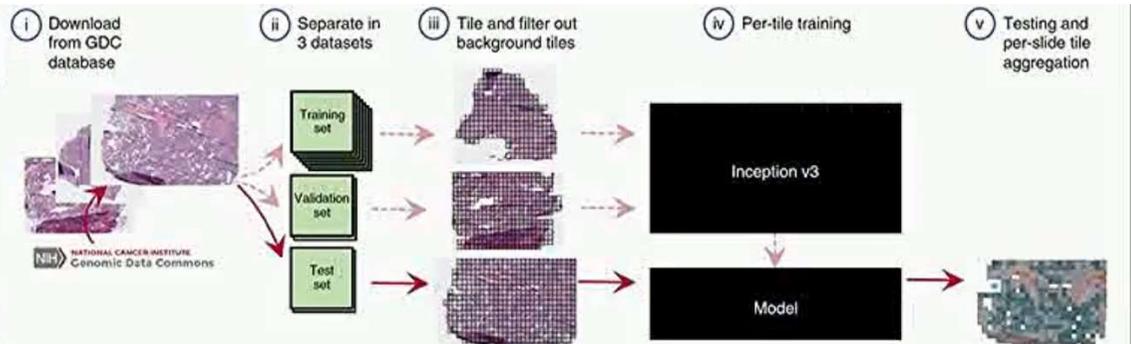
Visual inspection of histopathology slides is one of the main methods used by pathologists to assess the stage, type and subtype of lung tumors. Adenocarcinoma (LUAD) and squamous cell carcinoma (LUSC) are the most prevalent subtypes of lung cancer, and their distinction requires visual inspection by an experienced pathologist. In this study, we trained a deep convolutional neural network (Inception v3) on whole-slide images obtained from The Cancer Genome Atlas to accurately and automatically classify them into LUAD, LUSC or normal lung tissue. The performance of our method is comparable to that of pathologists, with an average area under the curve (AUC) of 0.97. Our model was validated on independent datasets of frozen tissues, formalin-fixed paraffin-embedded tissues and biopsies. Furthermore, we trained the network to predict the ten most commonly mutated genes in LUAD. We found that six of them—STK11, EGFR, FAT1, SETBP1, KRAS and TP53—can be predicted from pathology images, with AUCs from 0.733 to 0.856 as measured on a held-out population. These findings suggest that deep-learning models can assist pathologists in the detection of cancer subtype or gene mutations. Our approach can be applied to any cancer type, and the code is available at <https://github.com/ncoudray/DeepPATH>.

According to the American Cancer Society and the Cancer Statistics Center (see URLs), over 150,000 patients with lung cancer succumb to the disease each year (154,050 expected for 2018), while another 200,000 new cases are diagnosed on a yearly basis (234,030 expected for 2018). It is one of the most widely spread cancers in the world because of not only smoking, but also exposure to toxic chemicals like radon, asbestos and arsenic. LUAD and LUSC are the two most prevalent types of non-small cell lung cancer, and each is associated with discrete treatment guidelines. In the absence of definitive histologic features, this important distinction can be challenging and time-consuming, and requires confirmatory immunohistochemical stains.

Classification of lung cancer type is a key diagnostic process because the available treatment options, including conventional chemotherapy and, more recently, targeted therapies, differ for LUAD and LUSC. Also, a LUAD diagnosis will prompt the search for molecular biomarkers and sensitizing mutations and thus has a great impact on treatment options<sup>1</sup>. For example, epidermal growth factor receptor (EGFR) mutations, present in about 20% of LUAD, and anaplastic lymphoma receptor tyrosine kinase (ALK) rearrangements, present in <5% of LUAD, currently have targeted therapies approved by the Food and Drug Administration (FDA)<sup>2</sup>. Mutations in other genes, such as KRAS and tumor protein P53 (TP53) are very common (about 25% and 50%, respectively) but have proven to be particularly challenging drug targets so far<sup>3</sup>. Lung biopsies are typically used to diagnose lung cancer

type and stage. Virtual microscopy of stained images of tissues is typically acquired at magnifications of 20× to 40×, generating very large two-dimensional images (10,000 to >100,000 pixels in each dimension) that are oftentimes challenging to visually inspect in an exhaustive manner. Furthermore, accurate interpretation can be difficult, and the distinction between LUAD and LUSC is not always clear, particularly in poorly differentiated tumors; in this case, ancillary studies are recommended for accurate classification<sup>4,5</sup>. To assist experts, automatic analysis of lung cancer whole-slide images has been recently studied to predict survival outcomes<sup>6</sup> and classification<sup>7</sup>. For the latter, Yu et al.<sup>7</sup> combined conventional thresholding and image processing techniques with machine-learning methods, such as random forest classifiers, support vector machines (SVM) or Naive Bayes classifiers, achieving an AUC of ~0.85 in distinguishing normal from tumor slides, and ~0.75 in distinguishing LUAD from LUSC slides. More recently, deep learning was used for the classification of breast, bladder and lung tumors, achieving an AUC of 0.83 in classification of lung tumor types on tumor slides from The Cancer Genome Atlas (TCGA)<sup>8</sup>. Analysis of plasma DNA values was also shown to be a good predictor of the presence of non-small cell cancer, with an AUC of ~0.94 (ref. <sup>9</sup>) in distinguishing LUAD from LUSC, whereas the use of immunohistochemical markers yields an AUC of ~0.941<sup>10</sup>.

Here, we demonstrate how the field can further benefit from deep learning by presenting a strategy based on convolutional neural networks (CNNs) that not only outperforms methods in previously



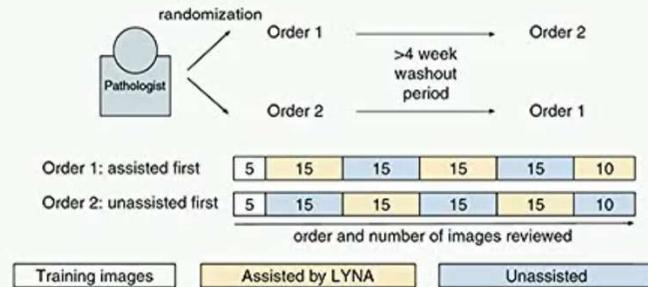
**Table 1 | AUC achieved by the network trained on mutations (with 95% CIs)**

Mutations	Per-tile AUC	Per-slide AUC after aggregation by...	
		... average predicted probability	... percentage of positive classified tiles
STK11	0.845 (0.838–0.852)	0.856 (0.709–0.964)	0.842 (0.683–0.967)
EGFR	0.754 (0.746–0.761)	0.826 (0.628–0.979)	0.782 (0.516–0.979)
SETBP1	0.785 (0.776–0.794)	0.775 (0.595–0.931)	0.752 (0.550–0.927)
TP53	0.674 (0.666–0.681)	0.760 (0.626–0.872)	0.754 (0.627–0.870)
FAT1	0.739 (0.732–0.746)	0.750 (0.512–0.940)	0.750 (0.491–0.946)
KRAS	0.814 (0.807–0.829)	0.733 (0.580–0.857)	0.716 (0.552–0.854)
KEAP1	0.684 (0.670–0.694)	0.675 (0.466–0.865)	0.659 (0.440–0.856)
LRP1B	0.640 (0.633–0.647)	0.656 (0.513–0.797)	0.657 (0.512–0.799)
FAT4	0.768 (0.760–0.775)	0.642 (0.470–0.799)	0.640 (0.440–0.856)
NF1	0.714 (0.704–0.724)	0.640 (0.419–0.845)	0.632 (0.405–0.845)

<sup>1</sup>Applied Bioinformatics Laboratories, New York University School of Medicine, New York, NY, USA. <sup>2</sup>Skirball Institute, Department of Cell Biology, New York University School of Medicine, New York, NY, USA. <sup>3</sup>Department of Pathology, New York University School of Medicine, New York, NY, USA. <sup>4</sup>School of Mechanical Engineering, National Technical University of Athens, Zografou, Greece. <sup>5</sup>Institute for Systems Genetics, New York University School of Medicine, New York, NY, USA. <sup>6</sup>Department of Biochemistry and Molecular Pharmacology, New York University School of Medicine, New York, NY, USA. <sup>7</sup>Center for Biopreparatory Research and Development, New York University, New York, NY, USA. <sup>8</sup>Department of Population Health and the Center for Healthcare Innovation and Delivery Science, New York University School of Medicine, New York, NY, USA. \*These authors contributed equally to this work.

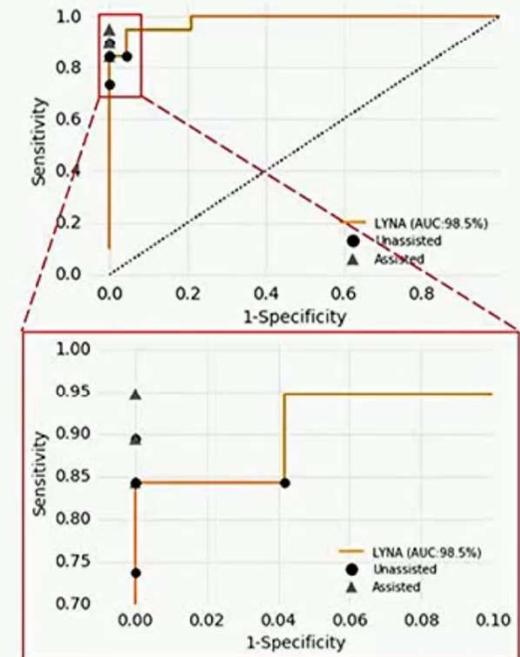
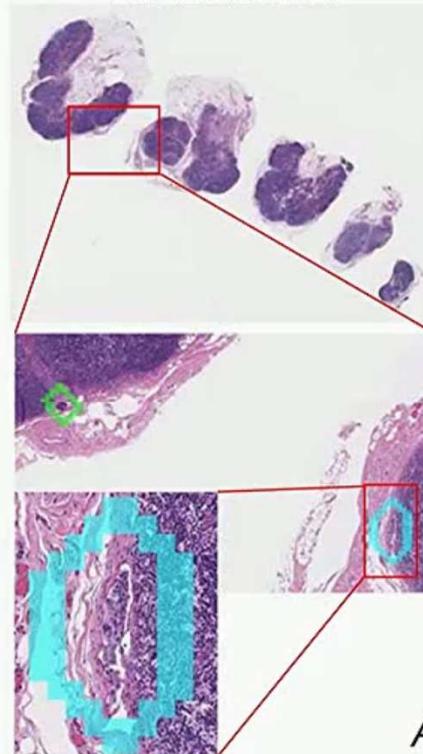
# Impact of Deep Learning Assistance on the Histopathologic Review of Lymph Nodes for Metastatic Breast Cancer

David F. Steiner, MD, PhD,\* Robert MacDonald, PhD,\* Yun Liu, PhD,\* Peter Truszkowski, MD,\*  
 Jason D. Hipp, MD, PhD, FCAP,\* Christopher Gammage, MS,\* Florence Thng, MS,†  
 Lily Peng, MD, PhD,\* and Martin C. Stumpe, PhD\*



6 pathologists/ 70 DSIs  
 AI + pathologist higher accuracy vs AI  
 or path  
 LYNA algorithm AUC .985  
 Marked increase in speed of review  
 for micrometastases: 61 v 116 secs

LYNA-assisted review



## Development and validation of a deep-learning algorithm for the detection of polyps during colonoscopy

Pu Wang<sup>1</sup>, Xiao Xiao<sup>2</sup>, Jeremy R. Glissen Brown<sup>3</sup>, Tyler M. Berzin<sup>4</sup>, Mengtian Tu<sup>1</sup>, Fei Xiong<sup>1</sup>, Xiao Hu<sup>1</sup>, Peixi Liu<sup>1</sup>, Yan Song<sup>1</sup>, Di Zhang<sup>1</sup>, Xue Yang<sup>1</sup>, Liangping Li<sup>1</sup>, Jiong He<sup>2</sup>, Xin Yi<sup>2</sup>, Jingjia Liu<sup>2</sup> and Xiaogang Liu<sup>1\*</sup>

The detection and removal of precancerous polyps via colonoscopy is the gold standard for the prevention of colon cancer. However, the detection rate of adenomatous polyps can vary significantly among endoscopists. Here, we show that a machine-learning algorithm can detect polyps in clinical colonoscopies, in real time and with high sensitivity and specificity. We developed the deep-learning algorithm by using data from 1,290 patients, and validated it on newly collected 27,113 colonoscopy images from 1,138 patients with at least one detected polyp (per-image-sensitivity, 94.38%; per-image-specificity, 95.92%; area under the receiver operating characteristic curve, 0.984), on a public database of 612 polyp-containing images (per-image-sensitivity, 88.24%), on 138 colonoscopy videos with histologically confirmed polyps (per-image-sensitivity of 91.64%; per-polyp-sensitivity, 100%), and on 54 unaltered full-range colonoscopy videos without polyps (per-image-specificity, 95.40%). By using a multi-threaded processing system, the algorithm can process at least 25 frames per second with a latency of  $76.80 \pm 5.60$  ms in real-time video analysis. The software may aid endoscopists while performing colonoscopies, and help assess differences in polyp and adenoma detection performance among endoscopists.

Colonoscopy is the gold-standard screening test for colorectal cancer<sup>1,2</sup>, one of the leading causes of cancer death in both the United States<sup>3</sup> and China<sup>4</sup>. Colonoscopy can reduce the risk of death from colorectal cancer through the detection of tumours at an earlier, more treatable stage as well as through the removal of precancerous adenomas<sup>5</sup>. Conversely, failure to detect adenomas may lead to the development of interval cancer. Evidence has shown that each 1.0% increase in adenoma detection rate (ADR) leads to a 3.0% decrease in the risk of interval colorectal cancer<sup>6</sup>.

Although more than 14 million colonoscopies are performed in the United States annually<sup>7</sup>, the adenoma miss rate (AMR) is estimated to be 6–27%. Certain polyps may be missed more frequently, including smaller polyps<sup>8,9</sup>, flat polyps<sup>10</sup> and polyps in the left colon<sup>11</sup>. There are two independent reasons why a polyp may be missed during colonoscopy: (i) it was never in the visual field or (ii) it was in the visual field but not recognized. Several hardware innovations have sought to address the first problem by improving visualization of the colonic lumen, for instance by providing a larger, panoramic camera view, or by flattening colonic folds using a distal-cap attachment. The problem of unrecognized polyps within the visual field has been more difficult to address<sup>12</sup>. Several studies have shown that observation of the video monitor by either nurses or gastroenterology trainees may increase polyp detection by up to 30%<sup>13,14</sup>. Ideally, a real-time automatic polyp-detection system could serve as a similarly effective second observer that could draw the endoscopist's eye, in real time, to concerning lesions, effectively creating an 'extra set of eyes' on all aspects of the video data with fidelity. Although automatic polyp detection in colonoscopy videos has been an active research topic for the past 20 years, performance levels close to that of the expert endoscopist<sup>15–21</sup> have not

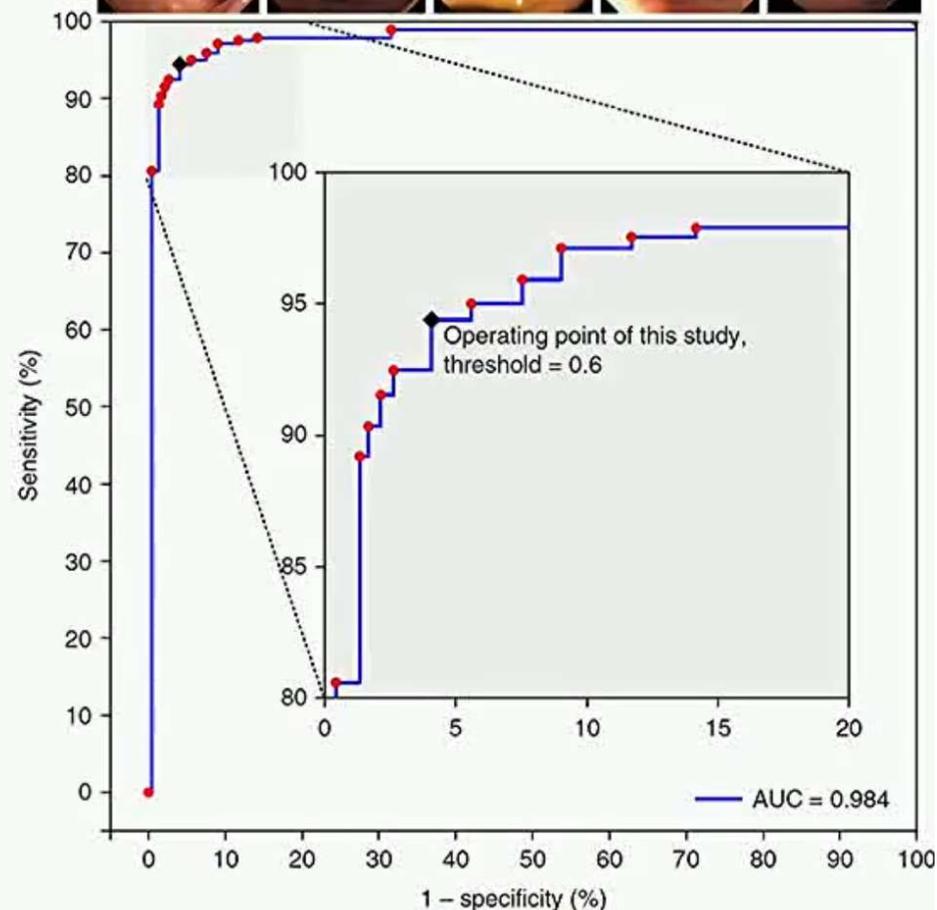
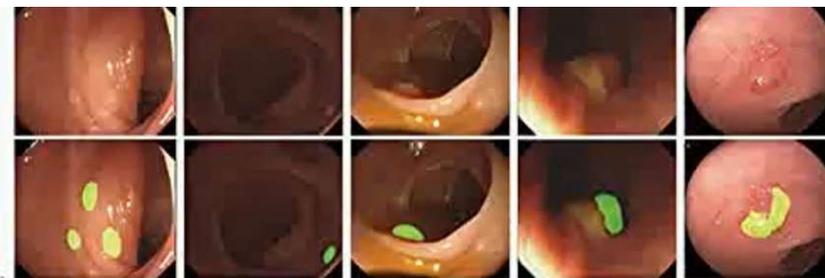
been achieved. Early work in automatic polyp detection has focused on applying deep-learning techniques to polyp detection, but most published works are small in scale, with small development and/or training validation sets<sup>22,23</sup>.

Here, we report the development and validation of a deep-learning algorithm, integrated with a multi-threaded processing system, for the automatic detection of polyps during colonoscopy. We validated the system in two image studies and two video studies. Each study contained two independent validation datasets.

### Results

We developed a deep-learning algorithm using 5,545 colonoscopy images from colonoscopy reports of 1,290 patients that underwent a colonoscopy examination in the Endoscopy Center of Sichuan Provincial People's Hospital between January 2007 and December 2015. Out of the 5,545 images used, 3,634 images contained polyps (65.54%) and 1,911 images did not contain polyps (34.46%). For algorithm training, experienced endoscopists annotated the presence of each polyp in all of the images in the development dataset. We validated the algorithm on four independent datasets. Datasets A and B were used for image analysis, and datasets C and D were used for video analysis.

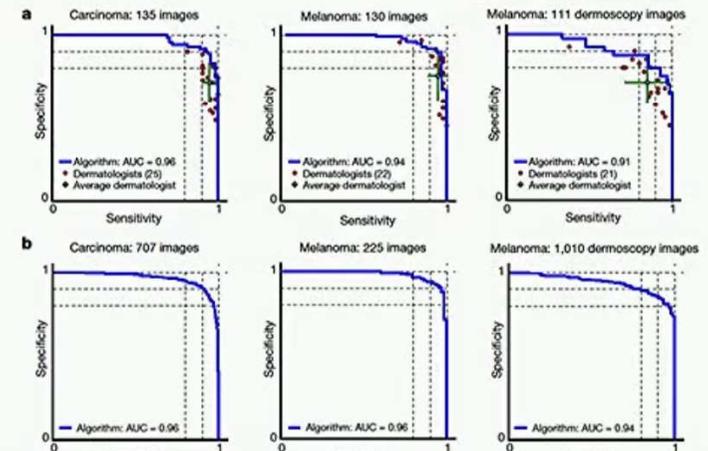
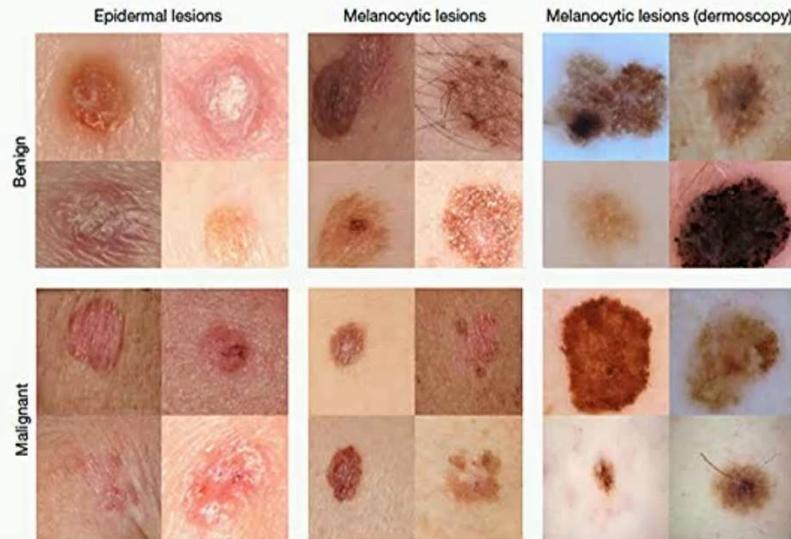
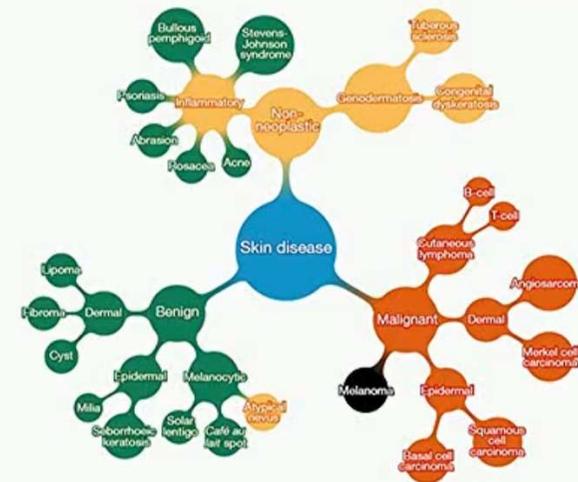
Dataset A contained 27,113 colonoscopy images from colonoscopy reports of 1,138 consecutive patients who underwent a colonoscopy examination in the Endoscopy Center of Sichuan Provincial People's Hospital between January and December 2016 and who were found to have at least one polyp. Out of the 27,113 images, 5,541 images contained polyps (20.44%) and 21,572 images did not contain polyps (79.56%). All polyps were confirmed histologically after biopsy. Dataset B is a public database (CVC-ClinicDB;



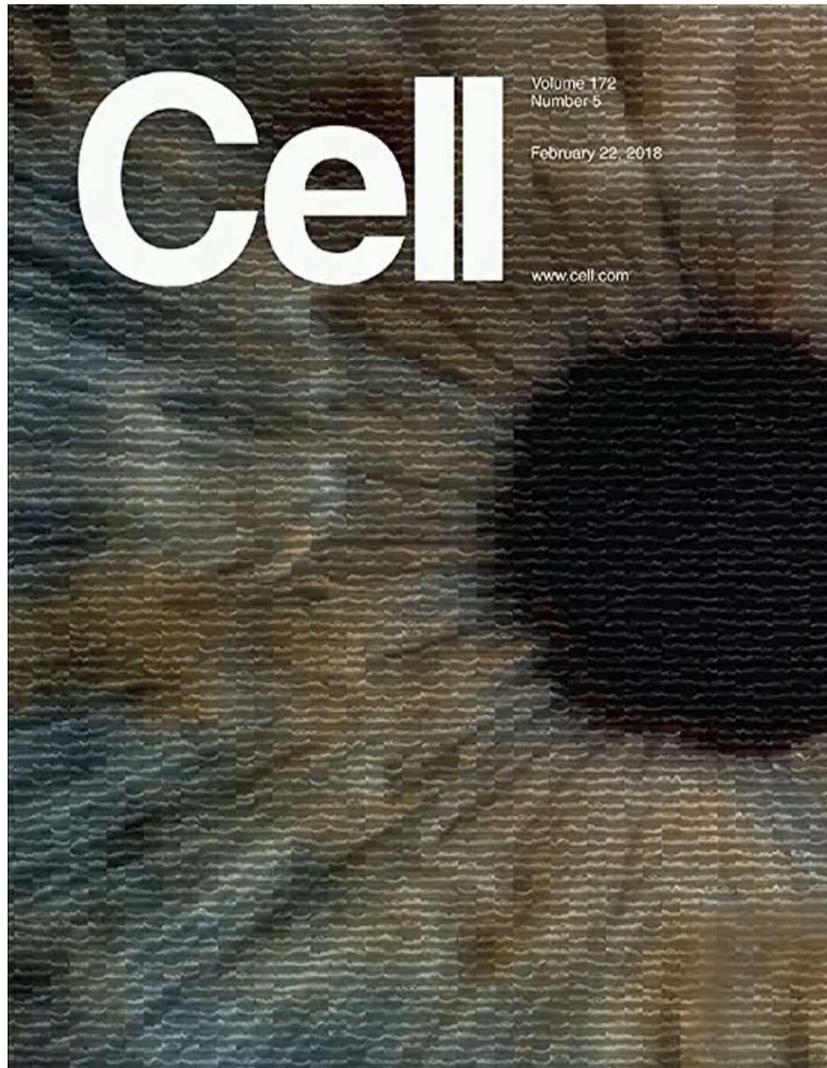
# Dermatologist-level classification of skin cancer with deep neural networks

nature

21 Board Certified Stanford Dermatologists  
 129, 450 images of 2,032 diseases  
 1.41 million AI training images



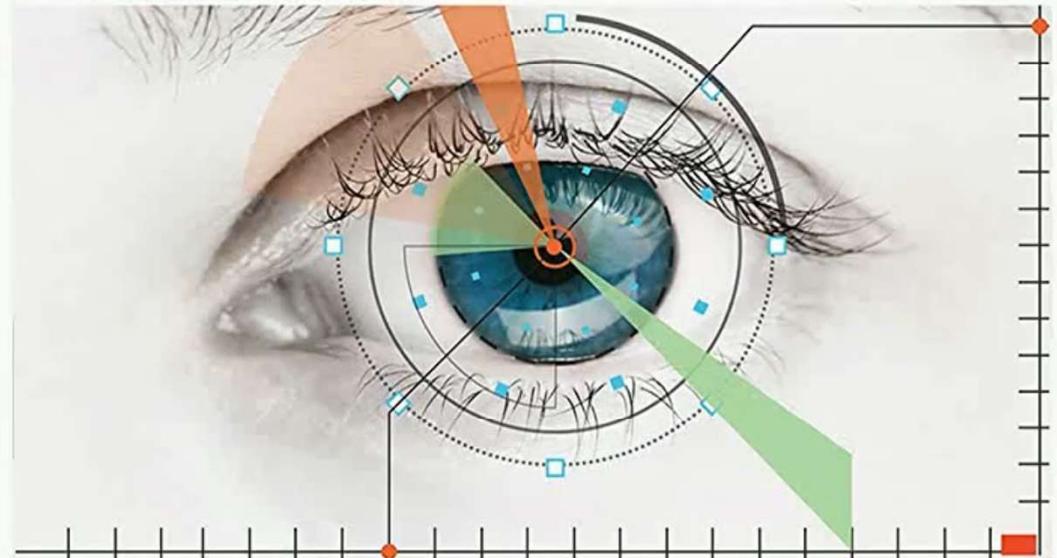
25 Jan 2017



## In the future, an AI may diagnose eye problems

New technology aims to speed medical care in places where specialists are scarce

BY MARIA TEMMING 8:00AM, MARCH 4, 2018



AI, M.D. New AI programs coming down the pipeline can examine eye images to diagnose blinding diseases and potentially other health problems.

ScienceNews

## Gender



Actual: Female  
Predicted: Female

# Prediction of cardiovascular risk factors from retinal fundus photographs via deep learning

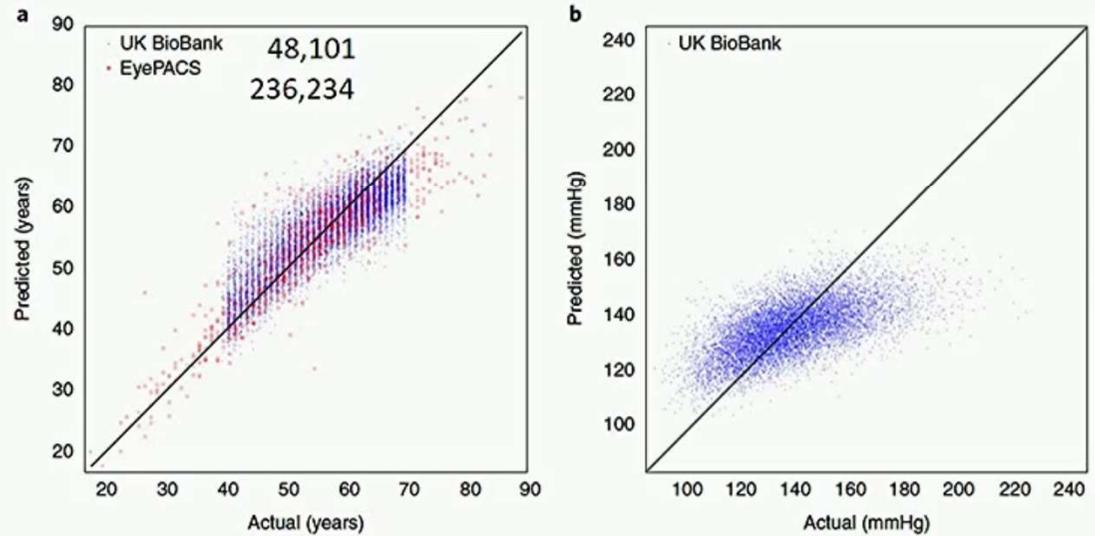
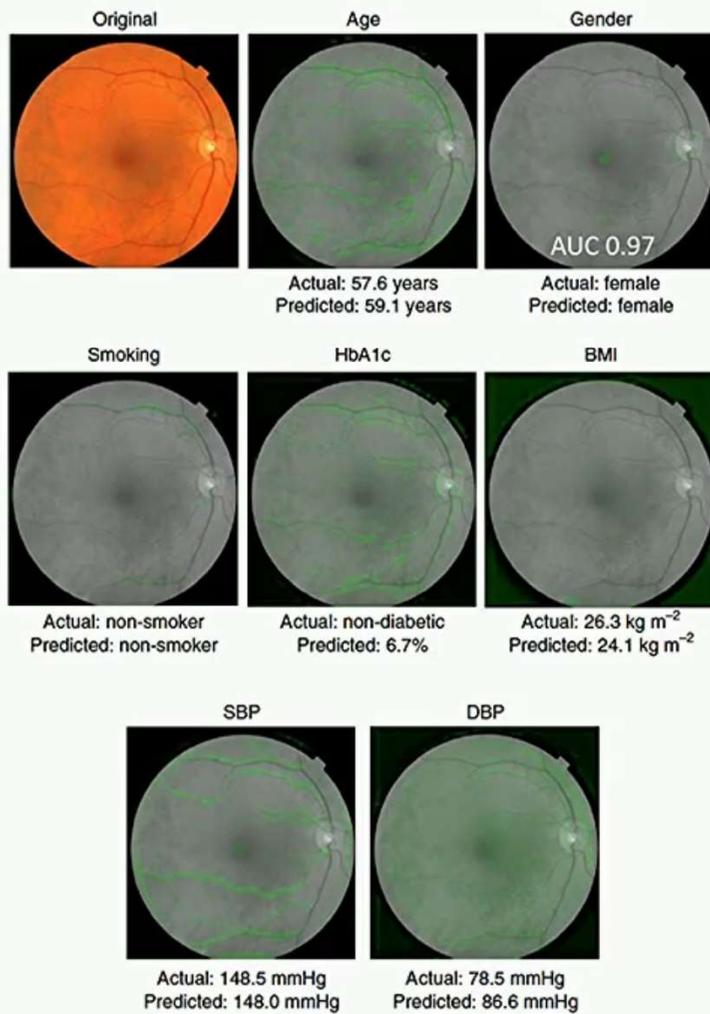
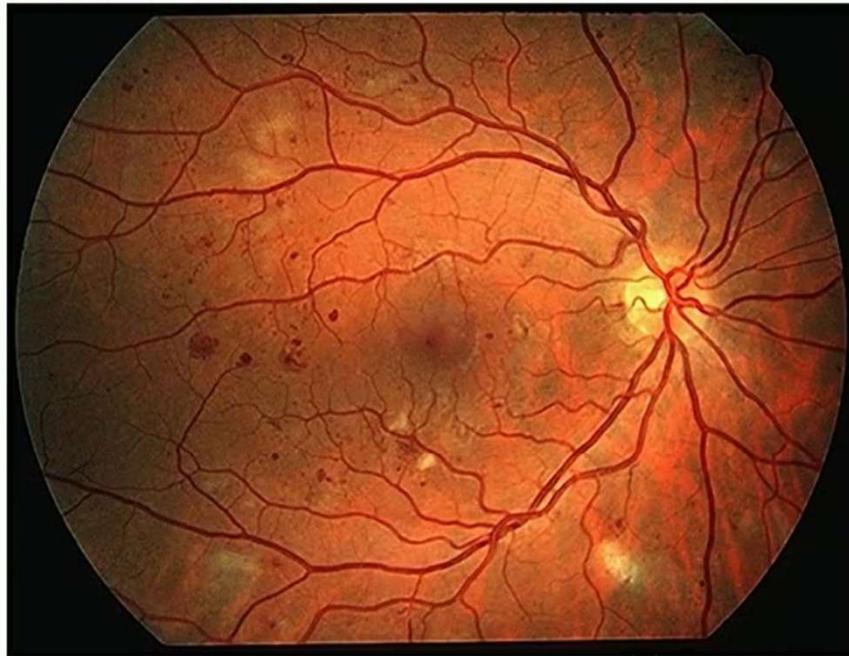


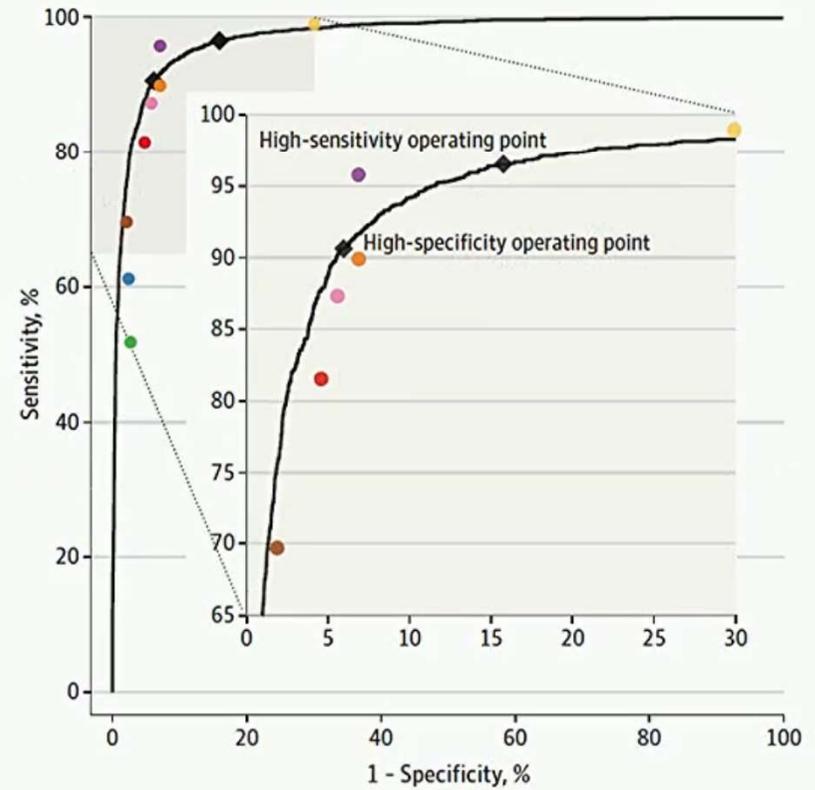
Fig. 1 | Predictions of age and SBP.

# Development and Validation of a Deep Learning Algorithm for Detection of Diabetic Retinopathy in Retinal Fundus Photographs



**JAMA** December 13, 2016

Figure 3. Validation Set Performance for All-Cause Referable Diabetic Retinopathy in the EyePACS-1 Data Set (9946 Images)



Performance of the algorithm (black curve) and ophthalmologists (colored circles)



## AI alone now making the diagnosis

By Rachel Z. Arndt

Modern Healthcare | October 8, 2018

ARTICLE [OPEN](#)

## Pivotal trial of an autonomous AI-based diagnostic system for detection of diabetic retinopathy in primary care offices

Michael D. Abràmoff<sup>1,2,3,4</sup>, Philip T. Lavin<sup>5</sup>, Michele Birch<sup>6</sup>, Nilay Shah<sup>7</sup> and James C. Folk<sup>1,2,3</sup>

### AI system diagnostic accuracy

	Point estimate	95% CI
Sensitivity	87.2%	81.8%–91.2%
Specificity	90.7%	88.3%–92.7%

EDITORIAL [OPEN](#)

## With an eye to AI and autonomous diagnosis

Pearse A. Keane<sup>1</sup> and Eric J. Topol<sup>2,3</sup>

<sup>1</sup>NIHR Biomedical Research Centre for Ophthalmology, Moorfields Eye Hospital NHS Foundation Trust and UCL Institute of Ophthalmology, London, UK; <sup>2</sup>Scripps Translational Science Institute,

**nature**  
npj | Digital Medicine 28 August 2018

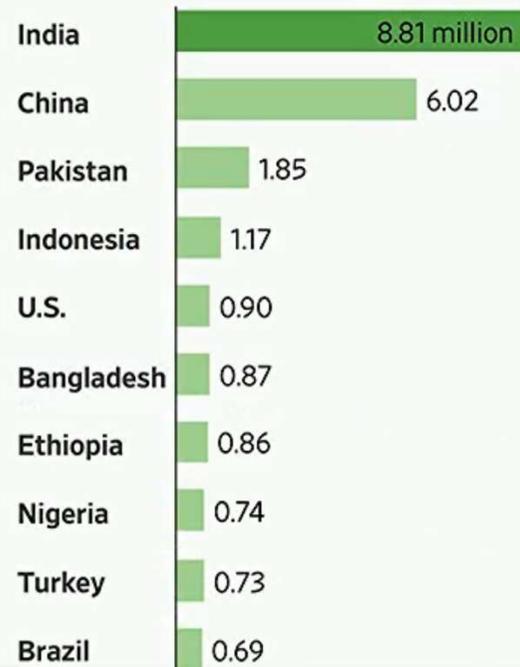
# Google's Effort to Prevent Blindness Shows AI Challenges

Company's AI can detect a condition that causes blindness in diabetes patients, but in rural India it doesn't always work



## Lacking Vision

India has the greatest number of people who are blind.



## Lacking Doctors

India has a shortfall of ophthalmologists.

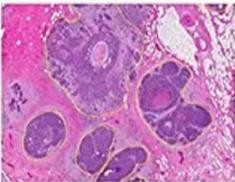
### Ophthalmologists per million people



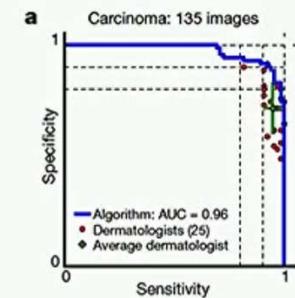
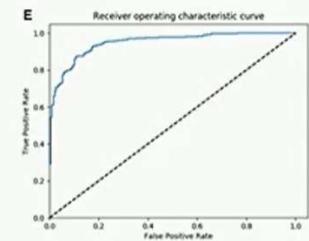
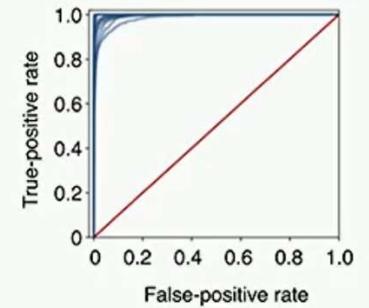
28 January

2019  
THE WALL STREET JOURNAL.

# How AI Will Transform Clinical Practice



Clinician Category	Data Interpretation
Radiologists	Scans
Pathologists	Slides
Dermatologists	Skin Lesions
Ophthalmologists	Eye Exams
Cardiologists	ECG, Echo
Psychiatrists	Ψ Status
Gastroenterologists	Scopes
Oncologists	Omics, Rx
Geneticists	Face, BAM file
Palliative Care	Predictions
All Doctors	Delete Keyboards
Nurses	Vital signs
Pharmacists	Drugs



## Digital Medicine Clinical Trials

Condition	Digital Intervention	Impact	Citation
Asthma	Inhaler Sensor + GPS for hot spots	Reduce rescue inhaler use by 78%; 48% more symptom-free days (Louisville Air)	Barrett, Health Affairs, April 2018
Hypertension	Smartphone app RCT	Improved medication adherence	Morawski, JAMA Internal Med 2018
Diabetes	Telemedicine RCT	Improved control of Type 2 diabetes	Wild et al, PLOS Medicine, 2016
Heart failure	Telemedicine RCT	Less hospital admission and mortality	Koehler, Lancet, 2018
Inflammatory Bowel Disease	Telemedicine RCT	Striking reduction in outpatient visits and hospital admissions	De Jong, Lancet 2017
Cancer	Smartphone app RCT	Improved survival in lung cancer	Denis, ASCO 2018
Headaches	Telemedicine RCT	As effective as traditional consultations	Muller, Neurology, 2017
Visual Impairment	Smartphone app RCT	Marked improvement of detection among school children in Kenya	Rono, Lancet Global Health, 2018
Insomnia	Digital CBT RCT	Major reduction in insomnia among patients with mental health conditions	Freeman, Lancet Psychiatry, 2017
Attention deficit disorder	Video game RCT	Significant improvement of attention performance in children and adolescents	Kollins, December 2017 and Proof of Concept PLOS One, 2018
Schizophrenia	Avatar CBT RCT	Significant reduction of hallucinations	Craig, Lancet Psychiatry 2017
Low Back Pain	Digital care RCT	Significant reduction of pain	Shebib, NPJ Nature Digital Medicine, 2019

CBT-coognitive behavioral therapy. RCT-randomized controlled trial



aidoc

iCAD

zebra  
MEDICAL VISION

MIRADA

BAYLABS

Neural Analytics

IDx

icometrix

IMAGEN

DENSITAS iz.ai

maxQ  
Artificial Intelligence

AliveCor

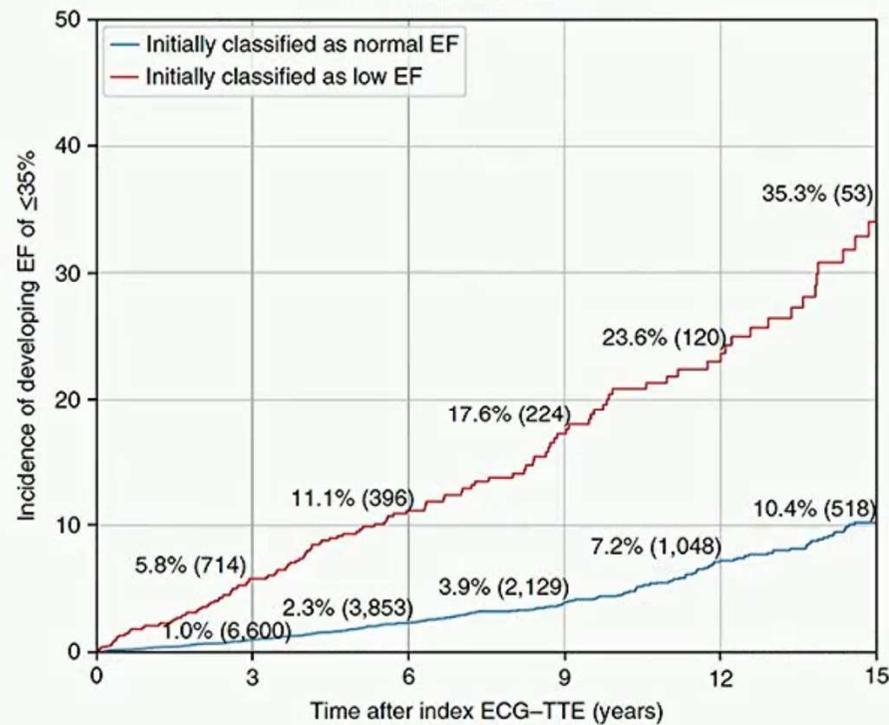
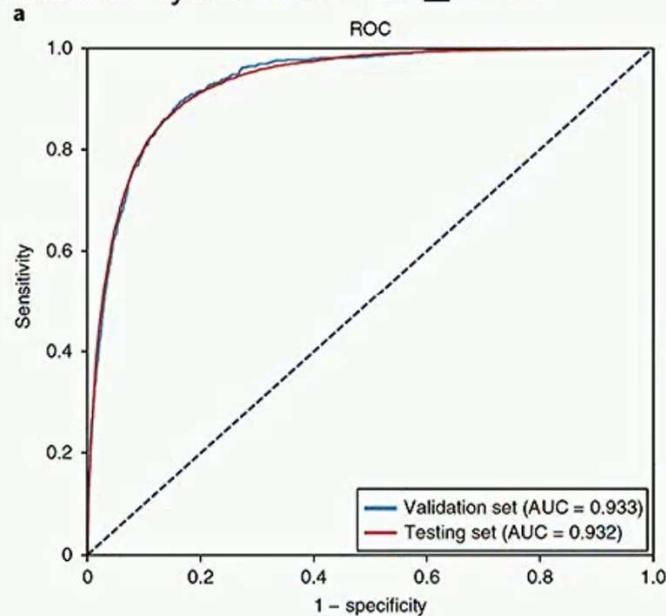
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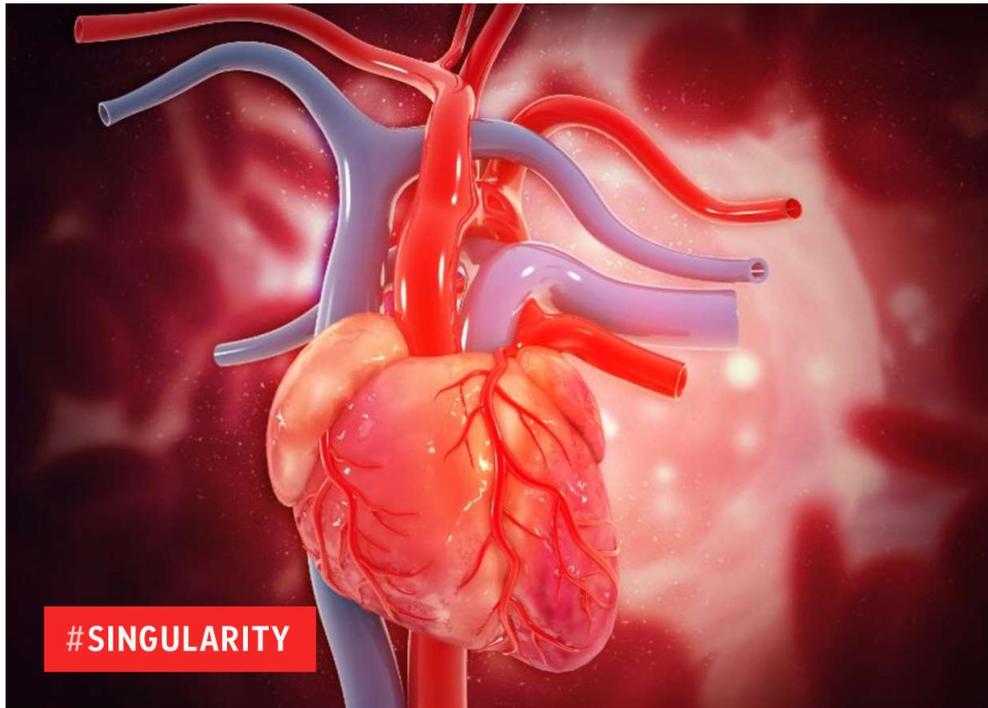
## FDA AI Approvals Are Accelerating

Company	FDA Approval	Indication
Apple	September 2018	Atrial fibrillation detection
Aidoc	August 2018	CT Brain bleed diagnosis
iCAD	August 2018	Breast density via mammography
Mirada Medical	July 2018	Radiation therapy planning
Zebra Medical	July 2018	Coronary calcium scoring
Bay Labs	June 2018	Echocardiogram EF determination
Neural Analytics	May 2018	Device for paramedic stroke diagnosis
Densasitas	April 2018	Breast density via mammography
IDx	April 2018	Diabetic retinopathy diagnosis
Icometrix	April 2018	MRI brain interpretation
Imagen	March 2018	X-ray wrist fracture diagnosis
Viz.ai	February 2018	CT Stroke diagnosis
Arterys	February 2018	Liver and lung cancer (MRI,CT) diagnosis
MaxQ-AI	January 2018	CT Brain bleed diagnosis
Alivecor	November 2017	Atrial fibrillation detection via Apple Watch
Arterys	January 2017	MRI heart interpretation

## Screening for cardiac contractile dysfunction using an artificial intelligence-enabled electrocardiogram

Training set 44,959 patients: ECG + Echo  
 Validation in 52,870 patients  
 Ventricular dysfunction = EF  $\leq$  35%





# Artificial Intelligence Detects Heart Failure From One Heartbeat With 100% Accuracy

Forbes · Nicholas Fearn

Doctors can detect heart failure from a single heartbeat with 100% accuracy using a new artificial intelligence-driven neural network. That's according to a recent study published in *Biomedical Signal Processing and Control*...

≡ Forbes



Unlike existing methods that are often time-consuming and inaccurate, their model combines advanced signal processing and machine learning tools on raw ECG signals to improve detection rates dramatically.

Dr [Sebastiano Massaro](#), associate professor of organizational neuroscience at the University of Surrey, said: “First, by assessing ECG directly, we confirm that with AI it is possible to accurately detect CHF looking beyond heart rate variability analysis. Thus, we have in general results that are more adherent to the real behavior of the affected heart.”

In another part of the experiment, a specific CNN model was used to improve the accuracy of CHF detection while taking into account comparable models.

“We focus on the detection of the pathology from one single heartbeat in excerpts of 5-minutes rather than in 24-hours recordings,” said Massaro.



## Artificial intelligence could create heart attack early warning system

sciencefocus.com · Jason Goodyer

The creators believe the system "could be saving lives within the next year." Technology developed using artificial intelligence could identify people...

Now, researchers at the University of Oxford have developed a new method of [identifying future risk of heart attack using machine learning](#) - an application of artificial intelligence that provides systems the ability to automatically learn and improve from experience without being explicitly programmed.

Dubbed the fat radiomic profile (FRP), the system looks for signs of inflammation, scarring and changes to the blood vessels that supply blood to the heart.

"Just because someone's scan of their coronary artery shows there's no narrowing, that does not mean they are safe from a heart attack," [Professor Charalambos Antoniades](#), BHF Senior Clinical Fellow at the University of Oxford.

"By harnessing the power of AI, we've developed a fingerprint to find 'bad' characteristics around people's arteries. This has huge potential to detect the early signs of disease, and to be able to take all preventative steps before a heart attack strikes, ultimately saving lives. We genuinely believe this

SmartECHO: 1D/2D/3D/4D  
Echocardiography to solve the  
workflow and quality demands of  
cardiac imaging



Cardio-testing and ECHO laboratories

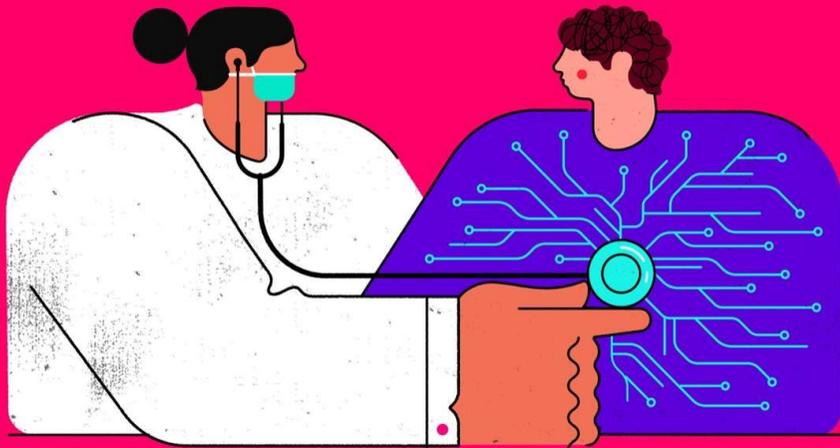
St Louis University Medical Center

St Louis, MO

Steven Smart MD, PhD

## How hospitals are using AI to save their sickest patients and curb 'alarm fatigue'

Early tests show artificial "assistants" can help doctors and nurses spot potentially deadly problems in time to take life-saving action.



— Early tests suggest artificial intelligence can improve patient care in hospitals' intensive care units while helping curb "alarm fatigue." Woody Harrington / for NBC News

In a traditional ICU, nurses respond to an alarm every 90 seconds, **two thirds of which turn out to be false alarms**, meaning they don't signal real danger. Some doctors and nurses have been known to tune out – **and even turn off** – alarms, a phenomenon called "alarm fatigue." In 2011, the FDA warned that alarm-related problems **contributed to more than 500 patient deaths from 2005 to 2008**.

Unlike a conventional monitoring system, AI is often able to predict problems hours in advance; doctors and nurses get a calm, text-message warning rather than having to respond to an urgent alarm signaling that a patient is already in trouble.

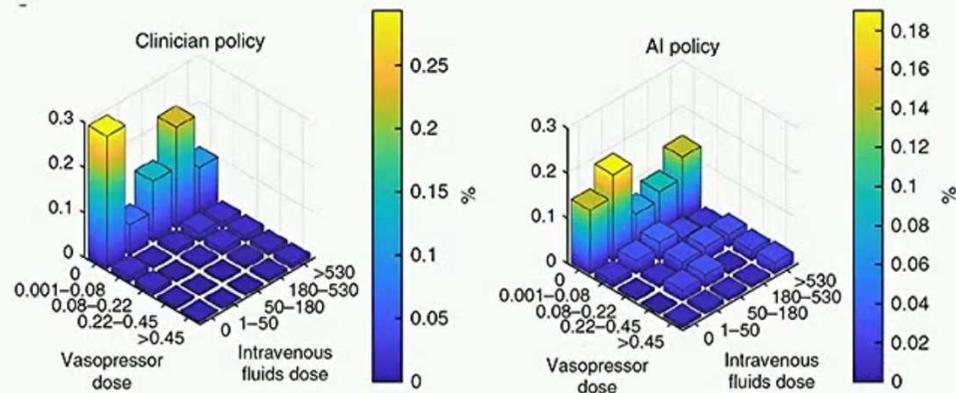
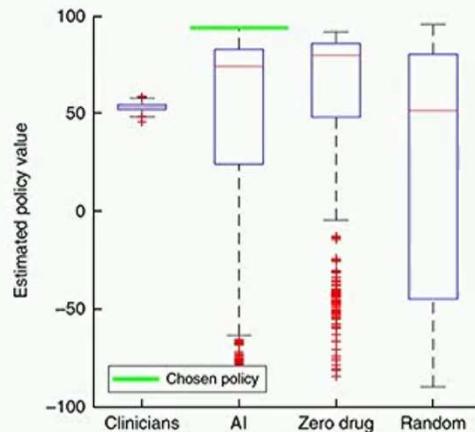
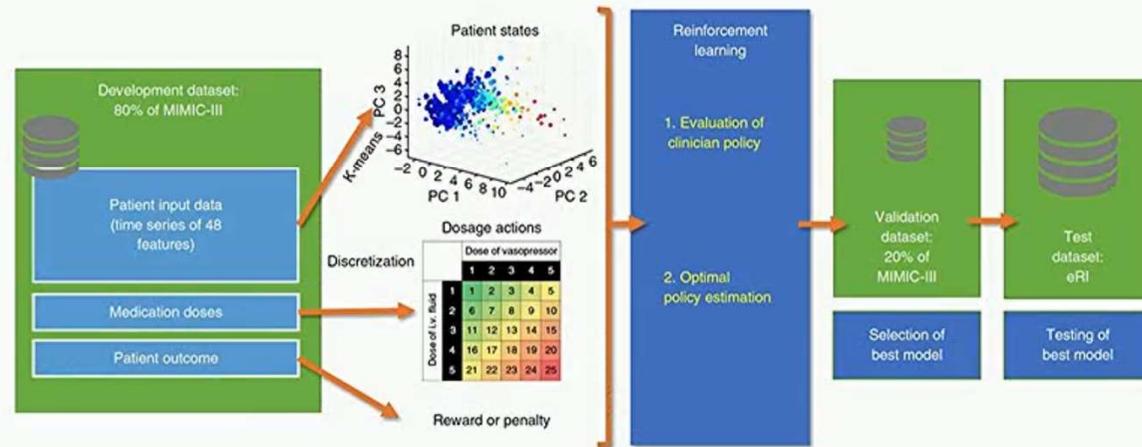
Barton said he received positive feedback from floor nurses during the AI study at UCSF. "The charge nurse of the floor will report that the number of false alarms has gone way down," he said. "It's making their work faster and more efficient."

17,803 admissions | 5 ICUs  
79,073 admissions | 128  
hospitals

Model for individualized patient  
AI: Better use of (less) IV fluids,  
(higher dose) vasopressors,  
medications

“AI Clinician is on average reliably  
higher than human clinicians”

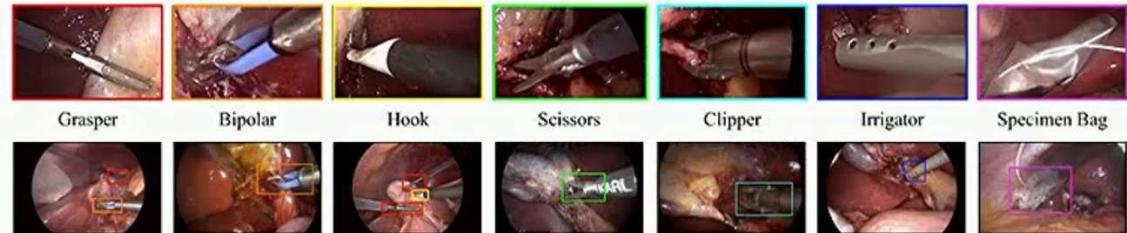
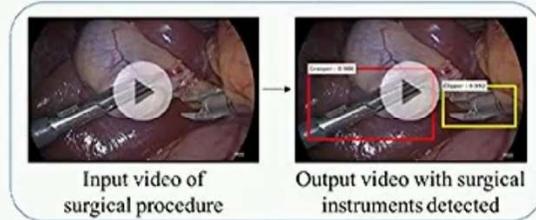
# The Artificial Intelligence Clinician learns optimal treatment strategies for sepsis in intensive care



22 October 2018

# Tool Detection and Operative Skill Assessment in Surgical Videos Using Region-Based Convolutional Neural Networks

Automatically detect surgical instruments

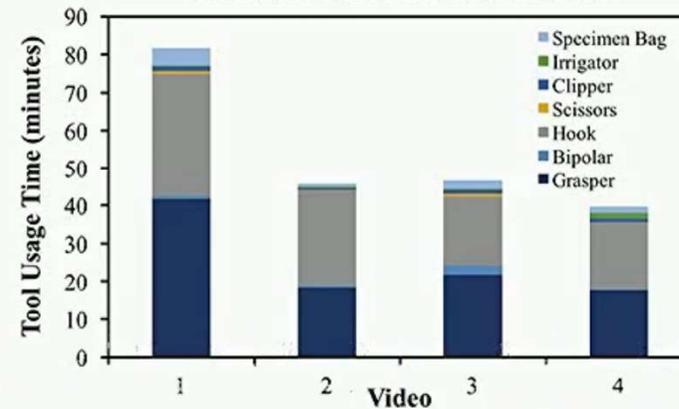


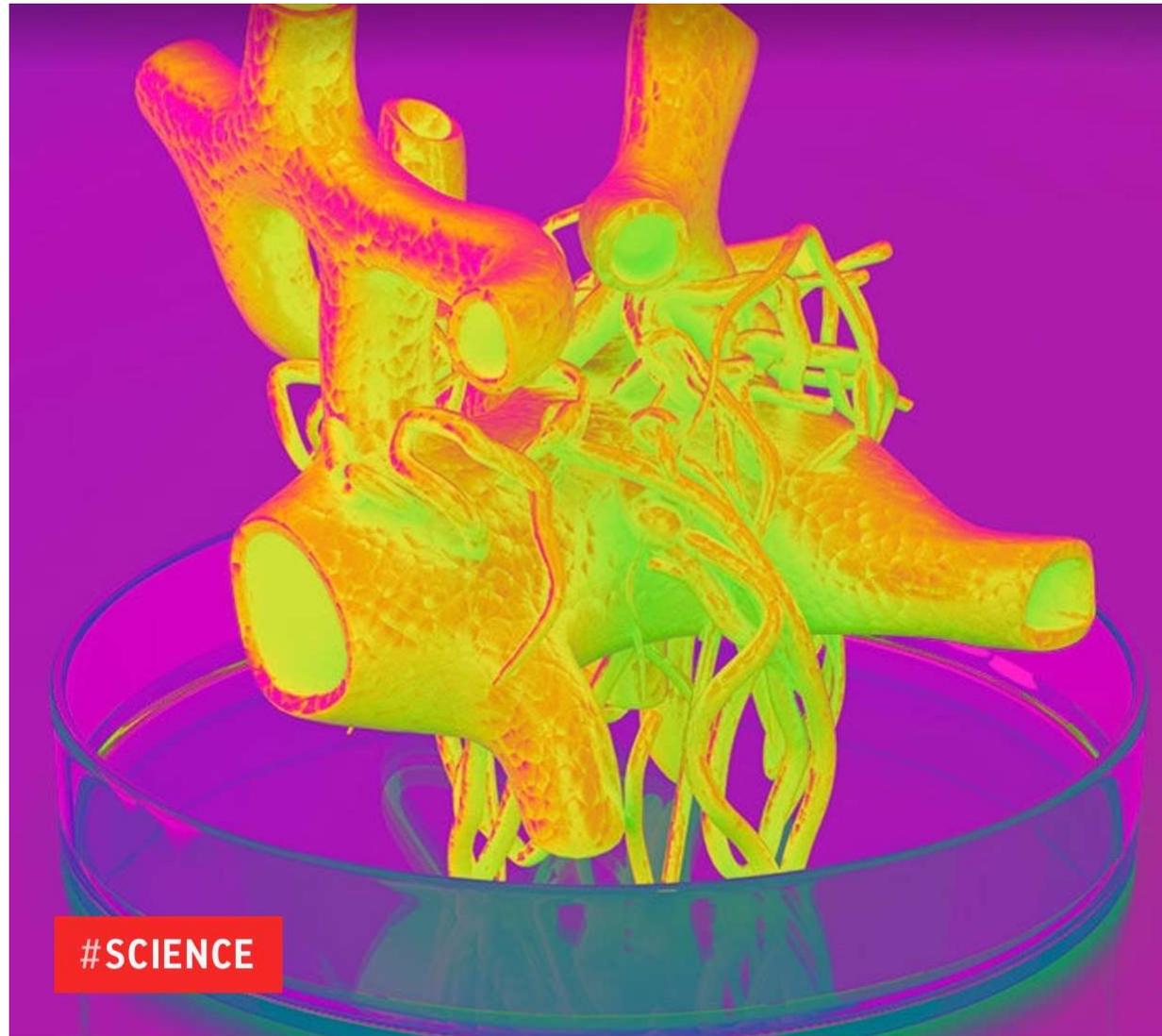
Extract metrics using tool detection results to assess operative skill



	Video 1	Video 2	Video 3	Video 4
<b>Depth Perception</b>	2.67	4.67	2.33	3.67
<b>Bimanual Dexterity</b>	3.00	4.67	2.00	3.33
<b>Efficiency</b>	2.00	4.67	2.33	3.00
<b>Tissue Handling</b>	2.33	4.67	2.67	3.33
<b>Total</b>	10.00	18.67	9.33	13.33

Total time each instrument is used





**Prepping for surgery with  
3D-printed organs may  
become commonplace**

Selected reports of machine and deep learning algorithms to predict clinical outcomes and related parameters

Prediction	N	AUC	Reference
In-hospital mortality, unplanned readmission, prolonged LOS, final discharge diagnosis	216,221	0.93* 0.75+ 0.85#	Rajkomar et al, Nature NPJ Digital Medicine, 2018
All-cause 3-12 month mortality	221,284	0.93^	Avati et al, arXiv, 2017
Readmission	1,068	0.78	Shameer et al, Pacific Symposium on Biocomputing, 2017
Sepsis	230,936	0.67	Hornig et al, PLOS One, 2017
Septic shock	16,234	0.83	Henry et al, Science, 2015
Severe sepsis	203,000	0.85@	Culliton et al, arXiv, 2017
C. Difficile infection	256,732	0.82++	Oh et al, Infection Control and Epidemiology, 2018
Developing diseases	704,587	range	Miotto et al, Scientific Reports, 2018
Diagnosis	18,590	0.96	Yang et al, Scientific Reports, 2018
Dementia	76,367	0.91	Cleret de Langavant et al, J Internet Med Res 2018
Alzheimer's Disease (+ amyloid imaging)	273	0.91	Mathotaarachchi et al, Neurobiology of Aging, 2017
Mortality after cancer chemotherapy	26,946	0.94	Elfiky et al, JAMA Open, 2018
Disease onset for 133 conditions	298,000	range	Razavian et al, arXiv, 2016
Suicide	5,543	0.84	Walsh et al, Clinical Psychological Science, 2017

# nature biomedical engineering

Explainable AI predicts blood-oxygen levels during anaesthesia

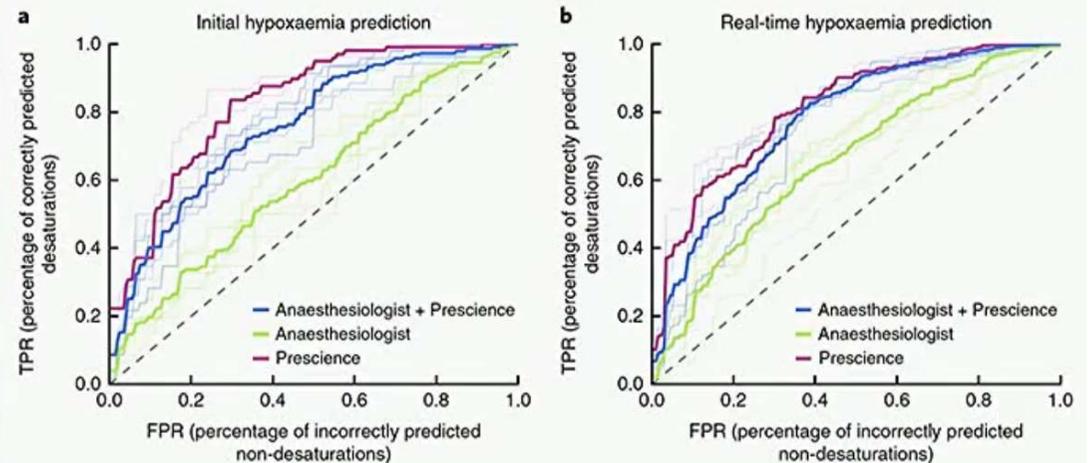
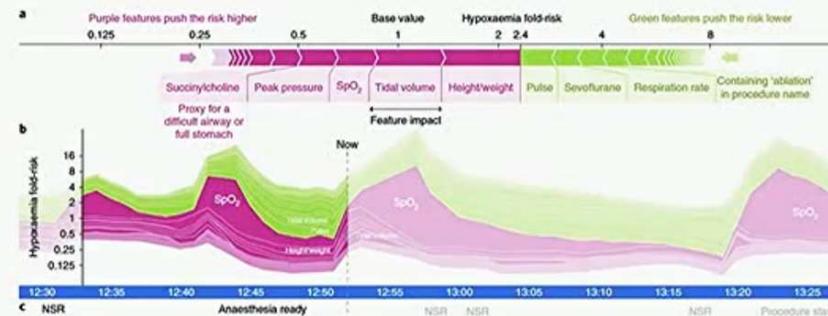
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<https://doi.org/10.1038/s41551-018-0304-0>

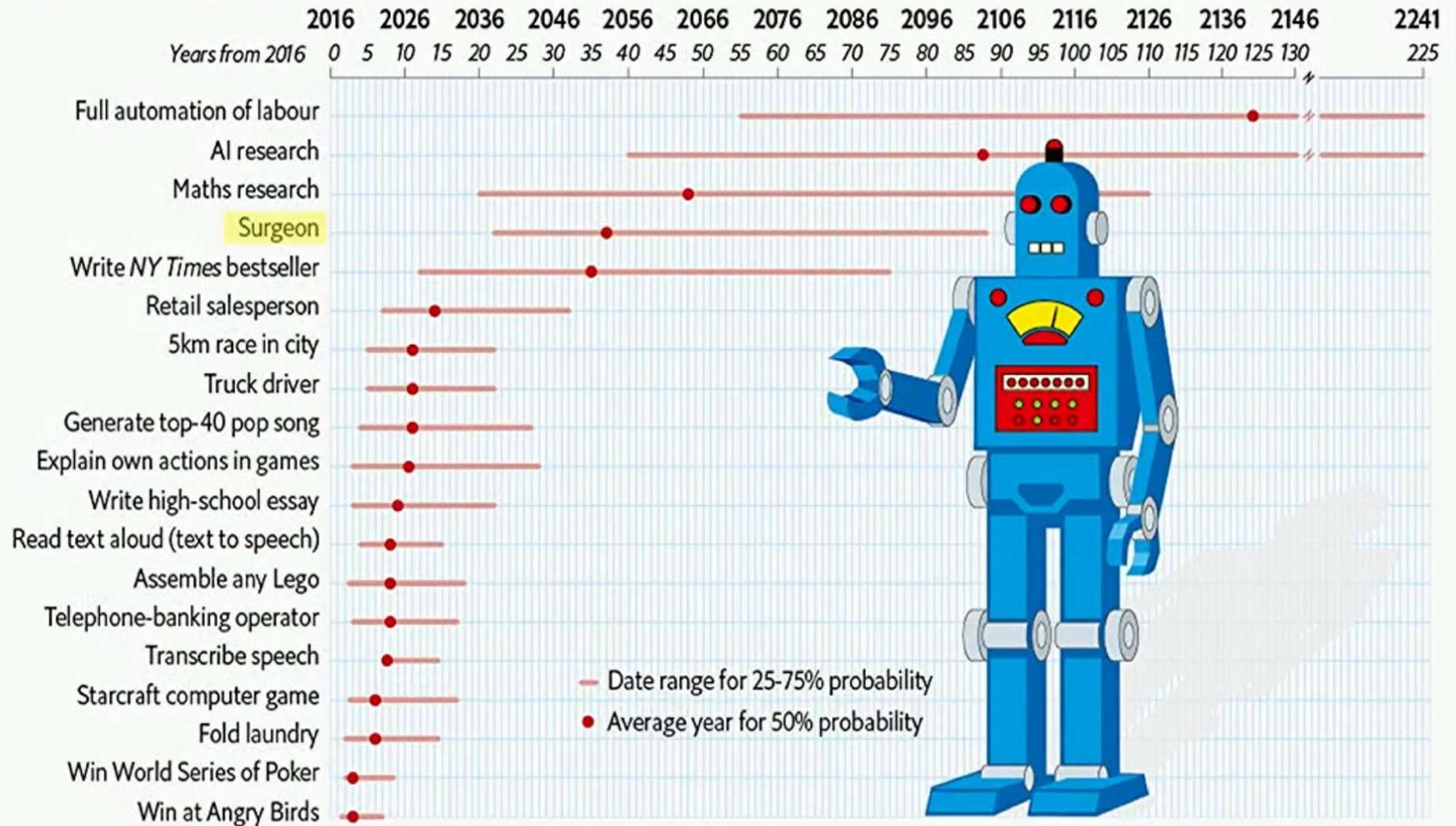
## Explainable machine-learning predictions for the prevention of hypoxaemia during surgery

Scott M. Lundberg<sup>1</sup>, Bala Nair<sup>2,3,4</sup>, Monica S. Vavilala<sup>2,3,4</sup>, Mayumi Horibe<sup>5</sup>, Michael J. Eisses<sup>2,6</sup>, Trevor Adams<sup>2,6</sup>, David E. Liston<sup>2,6</sup>, Daniel King-Wai Low<sup>2,6</sup>, Shu-Fang Newman<sup>2,3</sup>, Jerry Kim<sup>2,6</sup> and Su-In Lee<sup>1\*</sup>



# Man v machine

Predicted year machines will match human performance



Sources: Oxford University: Yale University

Our most valuable intelligence isn't artificial.



Together we th



# 31 GIORNATE CARDIOLOGICHE TORINESI

TURIN  
October  
24<sup>th</sup>-26<sup>th</sup>  
2019

# THANK YOU

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