Translating Artificial Intelligence Into Clinical Care

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Artificial intelligence has become a frequent topic in the news cycle, with reports of breakthroughs in speech recognition, computer vision, and textual understanding that have made

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their way into a bevy of products and services that are used every day. In contrast, clinical care has yet to reach the much lower bar of automating health care information transactions in the form

of electronic health records. Medical leaders in the 1960s and 1970s were already speculating about the opportunities to bring automated inference methods to patient care,¹ but the methods and data had not yet reached the critical mass needed to achieve those goals.

The intellectual roots of "deep learning," which power the commodity and consumer implementations of presentday artificial intelligence, were planted even earlier in the 1940s and 1950s with the development of "artificial neural network" algorithms.^{2,3} These algorithms, as their name suggests, are very loosely based on the way in which the brain's web of neurons adaptively becomes rewired in response to external stimuli to perform learning and pattern recognition. Even though these methods have had many success stories over the past 70 years, their performance and adoption in medicine in the past 5 years has seen a quantum leap. The catalyzing event occurred in 2012 when a team of researchers from the <mark>University of Toronto</mark> reduced the error rate in half on a well-known computer vision challenge using a deep learning algorithm.⁴ This work rapidly accelerated research and development in deep learning and propelled the field forward at a staggering pace. With the increased availability of digital clinical data, it remains to be seen how these deep learning models might be applied to the medical domain.

In this issue of *JAMA*, Gulshan and colleagues⁵ present findings from a study evaluating the use of deep learning for detection of diabetic retinopathy and macular edema. To build their model, the authors collected 128 175 annotated images from the EyePACs database. Each image was rated by 3 to 7 clinicians for referable diabetic retinopathy, diabetic macular edema, and overall image quality. Each rater was selected from a panel of 54 board-certified ophthalmologists and senior ophthalmology residents. Using this data set, the algorithm learned to predict the consensus grade of the raters along each clinical attribute: referable diabetic retinopathy, diabetic macular edema, and image quality. To validate their algorithm, the authors assessed its performance on 2 separate and nonoverlapping data sets consisting of 9963 and 1748 images. On the validation data, the algorithm had high sensitivity and specificity. Only one of these values (sensitivity on the second validation data set) failed to be superior at a statistically significant level. The other performance metrics (eg, area under the receiver operating characteristic curve, negative predictive value, positive predictive value) were likewise impressive, giving the authors confidence that this algorithm could be of clinical utility.

This work closely mirrors a recent "Kaggle" contest in which 661 teams competed to build an algorithm to predict the grade of diabetic retinopathy, albeit on a smaller data set with fewer grades per image. Kaggle is a website that hosts machine learning and data science contests. Companies and researchers can post their data to Kaggle and have contestants from around the world build predictive models. In the diabetic retinopathy contest, nearly all of the top teams used some form of deep learning and had little to no knowledge of the eye or ophthalmology. The first-place team⁶ and secondplace team⁷ both used standard deep learning models and were data science practitioners, not medical professionals. Gulshan et al correctly pointed out that a prerequisite for a successful deep learning model is access to a large database of images with high-quality annotations. Accordingly, the investigators increased both the number of images available and the number of ratings per image, which allowed them to improve on the existing state of the art with respect to both Kaggle and the existing scientific literature.

To build their algorithm, Gulshan et al leveraged a workhorse model in deep learning known as a convolutional neural network that has been critically important to recent advances in automatic image recognition. The convolutional neural network model used by the authors is known as the Inception-V3 network,⁸ which was developed by Google for entry in the Large Scale Visual Recognition Challenge, which it won in 2014. In this contest, known as ImageNet,⁹ researchers were given 1.2 million images that involve 1000 different categories that cover a wide variety of everyday objects, such as cats, dogs, automobiles, and different kinds of food. The goal of the contest was to build a classifier that could automatically recognize which object was present in an image and to identify which region of the image contained the object. This challenge was broad so that it covered many types of objects that a computer vision system could encounter in the real world.

As a result of this contest, several techniques¹⁰⁻¹² have been pioneered that improved the accuracy of these models immensely. As with the study by Gulshan et al, these improvements are beginning to trickle into other areas of computer vision, including medical image processing. For example, Gulshan et al not only used the same network that was originally built for ImageNet, they also used that network configuration to initialize their model for this study. This is often known as "transfer learning" and occurs when a network trained for one task (eg, ImageNet recognition) is used to bootstrap a network to be used for a different task (eg, detection of diabetic retinopathy). Gulshan et al observed a boost in performance when they used the parameters learned on ImageNet to initialize their model, thus demonstrating how progress in one domain can be used to accelerate progress in another.

Stepping back, one can consider how these results might affect medicine and, in particular, areas of medicine that involve the analysis of images such as pathology, radiology, and dermatology.¹³ It seems likely that these algorithms will reshape specific aspects of these specialties as more algorithms are developed to address a wider range of medical imaging tasks. Because these algorithms are by their nature standardized, repeatable, and scalable, they can be deployed to analyze a large number of images in hospitals around the world once an algorithm has been developed and validated, enabling clinicians to focus on other aspects of their practice.

A simple cost-benefit analysis reveals some interesting implications. Once a model has been "trained," it can be deployed on a relatively modest budget. Deep learning uses a specialized type of computer chip known as a graphics processing unit to process data at high speeds. A modern graphics processing unit costing approximately \$1000 can be added to most existing computer systems with little difficulty and can process about 3000 images per second¹⁴ depending on the complexity of the underlying deep learning model. This translates to an image processing capacity of almost 260 million images per day (because these devices can work around the clock), all for the cost of approximately \$1000. How will practice and clinical training adapt to refocus if initial screening of images is delegated to a machine with a learning algorithm? How will these capabilities mesh with current regulatory and reimbursement policies, or will these have to be modified?

Finally, the commercial efforts to push this technology into clinical care are becoming apparent, as several companies have begun to translate these research advancements to commercial applications. For example, one company is using deep learning models to improve cancer detection,¹⁵ while another company uses deep learning to read radiology images.¹⁶ Outside of imaging, other companies using artificial intelligence have started to help manage care, predict patient out-<mark>comes</mark>, or <mark>monitor</mark> patients through <mark>wearable</mark> devices</mark>, all in an attempt improve health care delivery. Given that artificial intelligence has a <mark>50-year history</mark> of promising to revolutionize medicine and failing to do so, it is important to avoid overinterpreting these new results. However, given the rapid and impressive progress in other areas of artificial intelligence, along with results such as those presented by Gulshan et al, there are valid reasons to remain cautiously optimistic that the time could now be right for artificial intelligence to transform the clinic into a much higher-capacity and lower-cost information processing care service.

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Published Online: November 29, 2016. doi:10.1001/jama.2016.17217

Conflict of Interest Disclosures: The authors have completed and submitted the ICMJE Form for Disclosure of Potential Conflicts of Interest and none were reported.

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VIEWPOINT

Adapting to Artificial Intelligence Radiologists and Pathologists as Information Specialists

INNOVATIONS IN HEALTH CARE DELIVERY

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Artificial intelligence-the mimicking of human cogni-This progress in imaging has changed the work of tion by computers—was once a fable in science fiction but is becoming reality in medicine. The combination of big data and artificial intelligence, referred to by some as the fourth industrial revolution,¹ will change radiology and pathology along with other medical specialties. Although reports of radiologists and pathologists being replaced by computers seem exaggerated,² these specialties must plan strategically for a future in which artificial intelligence is part of the health care workforce.

Radiologists have always revered machines and technology. In 1960, Lusted predicted "an electronic scannercomputer to examine chest photofluorograms, to separate the clearly normal chest films from the abnormal chest films."³ Lusted further suggested that "the abnormal chest films would be marked for later study by the radiologists."³ Lusted's intuitions were prescient: interpreting radiographs is pattern recognition; computers can recognize patterns and may be helpful because some roentgenographic analyses can be automated.

Nearly 60 years after Lusted's prediction, Enlitic, a technology company in Silicon Valley, inputted images of normal radiographs and radiographs with fractures into a computerized database.⁴ Using deep learning, a refined version of artificial neural networks, the

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computer developed rules that not only identified radiographs with fractures but highlighted the fractures. The computer received the image data rather than rules for their interpretation. The computer was not programmed regarding what to detect but developed algorithms necessary for fracture detection using deep learning.⁴ Deep learning is an autodidact–like an outstanding radiology resident, the more images it analyzes, the better it gets. The IBM prototype for artificial intelligence, Watson, can identify pulmonary embolism on computed tomography (CT) and detect abnormal wall motion on echocardiography.⁵ Watson has a boundless capacity for learning-and now has 30 billion images to review after IBM acquired Merge. Watson may become the equivalent of a general radiologist with super-specialist skills in every domain—a radiologist's alter ego and nemesis.

radiologists. Radiology, once confined to projectional images, such as chest radiographs, has become more complex and data rich. Cross-sectional imaging such as CT and magnetic resonance, by showing anatomy with greater clarity, has made diagnosis simpler in many instances; for example, a ruptured aneurysm is inferred on a chest radiograph but actually seen on CT. However, this has come at a price-the amount of data has increased markedly. For example, a radiologist typically views 4000 images in a CT scan of multiple body parts ("pan <mark>scan") in patients with multiple trauma.</mark> The abundance of data has changed how radiologists interpret images; from pattern recognition, with clinical context, to searching for needles in haystacks; from inference to detection. The radiologist, once a maestro with a chest radiograph, is now often visually fatigued searching for an occult fracture in a pan scan.

The amount of data continues to increase in imaging, both extractable by the human eye and extractable only by software.⁶ Thus, radiology has moved from a subjective perceptual skill to an objective science. Data have empowered radiologists but also challenged them computationally because of their abundance and complexity. This has paved the way for the role of computers,

> which extract fine information about tissues invisible to the human eye and process those data quickly and accurately.

> How should the changes in imaging, coupled with artificial intelligence, further change the work of radiologists? To avoid being replaced by computers, ra-

diologists must allow themselves to be displaced by computers. While some radiographic analyses can be automated, others cannot. Radiologists should identify cognitively simple tasks that could be addressed by artificial intelligence, such as screening for lung cancer on CT. This involves detecting, measuring, and characterizing a lung nodule, the management of which is standardized.⁷ A radiology residency or a medical degree is not needed to detect lung nodules. Likewise, radiologists are overtrained to interpret portable chest radiographs obtained in the intensive care unit to confirm that support lines are in proper position. These studies are not challenging and may be ideal for automation and delegation to artificial intelligence.

The primary purpose of radiologists is the provision of medical information; the image is only a means to information. Radiologists are more aptly considered "information specialists" specializing in medical imaging. This is similar to pathologists, who are also information specialists. Pathologists and radiologists are fundamentally similar because both extract medical information from images.

Pathologists have embraced machines and technologies. Some tasks once performed manually by pathologists have been automated, such as cell counts, typing and screening of blood, and Papanicolaou tests, leaving pathologists with more complex tasks. Artificial intelligence can perform the more complex tasks of pathologists and, in some instances, with superior accuracy. A recent study showed that computers could predict the grade and stage of lung cancer better than pathologists.⁸ Even though such studies need larger-scale validation with more diverse tissue types, it is clear in both radiology and pathology that many tasks can be handled by artificial intelligence. To underscore the commonality between radiology and pathology, researchers using operant conditioning trained pigeons to spot abnormal calcifications on mammograms and detect breast cancer on histology.⁹

Because pathology and radiology have a similar past and a common destiny, perhaps these specialties should be **merged** into a single entity, the "information specialist," whose responsibility will not be so much to extract information from images and histology but to manage the information extracted by artificial intelligence in the clinical context of the patient.

The information specialist would not spend time inferring conditions between competing shadows on radiographs, scroll through hundreds of images looking for pulmonary embolus on CT, or examine slides for "orphan Annie"-shaped nuclei. Artificial intelligence could perform many such tasks. The information specialist would interpret the important data, advise on the added value of another diagnostic test, such as the need for additional imaging, anatomical pathology, or a laboratory test, and integrate information to guide clinicians. Radiologists and pathologists will still be the physician's physician.

Together, the information specialist and artificial intelligence could manage individuals and populations. If a single artificial intelligence unit could do the work of many radiologists, then a single information specialist could manage many units of artificial intelligence. This would truly scale the influence of radiologists and pathologists. If artificial intelligence becomes adept at screening for lung and breast cancer, it could screen populations faster than radiologists and at a fraction of cost. The information specialist could ensure that images are of sufficient quality and that artificial intelligence is yielding neither too many false-positive nor too many falsenegative results. The efficiency from the economies of scale because of artificial intelligence could benefit not just developed countries, such as the United States, but developing countries hampered by access to specialists. A single information specialist, with the help of artificial intelligence, could potentially manage screening for an entire town in Africa.

Information specialists should train in the traditional sciences of pathology and radiology. The training should take no longer than it presently takes because the trainee will not spend time mastering the pattern recognition required to become a competent radiologist or pathologist. Visual interpretation will be restricted to perceptual tasks that artificial intelligence cannot perform as well as humans. The trainee need only master enough medical physics to improve suboptimal quality of medical images. Information specialists should be taught Bayesian logic, statistics, and data science and be aware of other sources of information such as genomics and biometrics, insofar as they can integrate data from disparate sources with a patient's clinical condition.

There may be resistance to merging 2 distinct medical specialties, each of which has unique pedagogy, tradition, accreditation, and reimbursement. However, artificial intelligence will change these diagnostic fields. The merger is a natural fusion of human talent and artificial intelligence. United, radiologists and pathologists can thrive with the rise of artificial intelligence.

The history of automation in the broader economy has a reassuring message.¹ Jobs are not lost; rather, roles are redefined; humans are displaced to tasks needing a human element. Radiologists and pathologists need not fear artificial intelligence but rather must adapt incrementally to artificial intelligence, retaining their own services for cognitively challenging tasks. A unified discipline, information specialists would best be able to captain artificial intelligence and guide medical information to improve patient care.

ARTICLE INFORMATION

Published Online: November 29, 2016. doi:10.1001/jama.2016.17438

Conflict of Interest Disclosures: The authors have completed and submitted the ICMJE Form for Disclosure of Potential Conflicts of Interest. Dr Jha reports speaker fees from Toshiba Medical Systems. Dr Topol reports advisory fees from Google Inc and Apple. No other disclosures were reported.

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