

Y-LW reports personal fees from AstraZeneca, Eli Lilly, Roche, Pfizer, Bristol-Myers Squibb, Merck, and Boehringer Ingelheim; consulting or advisory role with AstraZeneca, Roche, Merck, and Boehringer Ingelheim; and research funding to Guangdong Lung Cancer Institute from Boehringer Ingelheim, AstraZeneca, and Roche. YL declares no competing interests.

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Machine versus man in skin cancer diagnosis

Skin examinations for diagnosis of skin cancer are facilitated by the use of dermatoscopy, a non-invasive skin imaging technique that amplifies features of pigmented skin lesions that are not easily discernable when examined by the naked eye. Since its introduction in the late 1980s, dermatoscopy has substantially increased diagnostic accuracy in the identification of melanoma. However, interpretation of dermatoscopic criteria remains a subjective procedure that requires specific and dedicated training. Several strategies based on artificial intelligence and machine learning, including deep learning networks, have been developed as aids for less experienced dermatologists or other physicians to classify dermatoscopic images into benign and malignant skin tumours, as well as more specific diagnostic categories,^{1–4} which in the long-term could revolutionise dermatology practice.

In *The Lancet Oncology*, Philipp Tschandl and colleagues⁵ compared the accuracy of diagnoses produced by human readers with that of machine-learning algorithms for dermatoscopic images of pigmented skin lesions. Most outcome measures were outperformed by the machine-learning algorithms with respect to human readers. The best machine-learning algorithms achieved a mean of 7.94 (95% CI 7.76–8.12; $p < 0.0001$) more correct diagnoses than the average human reader, and a mean of 6.65 (6.06–7.25; $p < 0.0001$) more correct diagnoses than expert human

readers with more than 10 years of experience.⁵ This robustly powered study provides a timely contribution to an increasingly exciting debate in the field of dermatology and focuses on the more general issue of artificial intelligence use in diagnostic medicine. Tschandl and colleagues have aggregated a hugely balanced amount of information across patient populations from different geographical areas and human readers with varying experience from a large variety of geographically, ethnically, and culturally diverse countries. Sets of images representative of various types of disease (benign and malignant) were analysed and coupled with a robust statistical model, resulting in sound acceptability. The comparison of a large series of different machine-learning algorithms, which were developed by 77 laboratories that participated in the International Skin Imaging Collaboration Forum 2018 challenge,⁶ is also novel.

Despite a marked effort to control for the intrinsic drawbacks of overfitting in machine learning, the authors underline that training too hard causes the predictive models to be too specific to the training set used.⁵ Overfitting emerges as a problem with models that fit to the training data too closely, thus limiting the generalisability of the model and leading to poor success for data that might seem similar, but are actually different. The training set in an overfitted model might generate a very high performance, which



Published Online
June 11, 2019
[http://dx.doi.org/10.1016/S1470-2045\(19\)30391-2](http://dx.doi.org/10.1016/S1470-2045(19)30391-2)
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is unfortunately associated with poor to mediocre performance on unseen data. Tschandl and colleagues⁵ also tested the machine-learning algorithms on images from sources different from those of the training set to properly quantify the overfitting issue.

Creating a gold standard of medical diagnosis is not a straightforward process. In a previous study by Tschandl and colleagues,⁷ more than 50% of lesions, including all malignant tumours, were analysed histopathologically. Identification of benign unexcised lesions included more than 1.5 years of follow-up of sequential dermatoscopic imaging without changes, and judgment on the basis of expert consensus in cases of benign, banal, non-melanocytic lesions. For conclusive classification, seven simplified classes of diseases were generated, with specific attention paid to avoiding ambiguous classifications, and cases with uncertain histopathological diagnoses were excluded.

Where and when to appropriately use artificial intelligence is a matter of initial, but already intense, debate. Although appreciated as a powerful tool, artificial intelligence does not seem ready to replace the refined cognitive process of integrating morphological observations within a clinical context. In a real-world situation, in addition to dermatoscopic images clinicians consider several other features, including anatomical site, age, sex, and clinical history and evolution. Machine learning and humans should also be compared in terms of turnaround time and cost-effectiveness before becoming a standard of care.

In certain high-need settings, such as low-income and middle-income countries or health-care systems with few human resources, artificial intelligence might bring additional value, particularly as a screening tool. Furthermore, artificial intelligence could also be useful for the development of clinical support tools for inexperienced physicians and general practitioners,

in view of an increasing demand for points of care for dermatology screening.⁸ However, in future, automated classifiers and artificial intelligence algorithms could be integrated into clinical dermatology practice for a more accurate and effective triage of lesions in the context of human-machine collaboration. Although there is little doubt that artificial intelligence and machine-learning algorithms for skin cancer diagnosis are already gaining a central role in dermatology research, it is possible to anticipate that their application to clinical practice will require serious and robust validation in large, prospective studies.

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We declare no competing interests.

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Neoadjuvant therapy for melanoma: is it ready for prime time?

In the past, advanced melanoma with nodal or distant metastases was almost universally fatal, with a median overall survival of less than 1 year with few effective therapies.¹ However, with the advent of treatment

with immune checkpoint blockade and BRAF-targeted therapy, among other strategies, a dramatic improvement has been seen in the survival of patients with stage IV disease, substantiating approval of such

Published Online
June 3, 2019
[http://dx.doi.org/10.1016/S1473-2045\(19\)30377-8](http://dx.doi.org/10.1016/S1473-2045(19)30377-8)

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