

## G OPEN ACCESS

**Citation:** Balagopalan A, Baldini I, Celi LA, Gichoya J, McCoy LG, Naumann T, et al. (2024) Machine learning for healthcare that matters: Reorienting from technical novelty to equitable impact. PLOS Digit Health 3(4): e0000474. https://doi.org/ 10.1371/journal.pdig.0000474

**Editor:** Omar Badawi, Telemedicine and Advanced Technology Research Center, UNITED STATES

Received: September 6, 2023

Accepted: February 18, 2024

Published: April 15, 2024

**Peer Review History:** PLOS recognizes the benefits of transparency in the peer review process; therefore, we enable the publication of all of the content of peer review and author responses alongside final, published articles. The editorial history of this article is available here: https://doi.org/10.1371/journal.pdig.0000474

**Copyright:** © 2024 Balagopalan et al. This is an open access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

**Data Availability Statement:** All data is contained within the manuscript.

**Funding:** A.B. was funded in part by an Amazon Science PhD Fellowship at the MIT Science Hub. I.

**RESEARCH ARTICLE** 

# Machine learning for healthcare that matters: Reorienting from technical novelty to equitable impact

Aparna Balagopalan<sup>1</sup>°, Ioana Baldini<sup>2</sup>°, Leo Anthony Celi<sup>3,4,5</sup>°, Judy Gichoya<sup>6</sup>°, Liam G. McCoy<sup>7</sup>°\*, Tristan Naumann<sup>8</sup>°, Uri Shalit<sup>9</sup>°, Mihaela van der Schaar<sup>10,11</sup>°, Kiri L. Wagstaff<sup>12</sup>°

 Department of Electrical Engineering and Computer Science, Massachusetts Institute of Technology; Cambridge, Massachusetts, United States of America, 2 IBM Research; Yorktown Heights, New York, United States of America, 3 Laboratory for Computational Physiology, Massachusetts Institute of Technology; Cambridge, Massachusetts, United States of America, 4 Division of Pulmonary, Critical Care and Sleep Medicine, Beth Israel Deaconess Medical Center; Boston, Massachusetts, United States of America,
Department of Biostatistics, Harvard T.H. Chan School of Public Health; Boston, Massachusetts, United States of America, 6 Department of Radiology and Imaging Sciences, School of Medicine, Emory University; Atlanta, Georgia, United States of America, 7 Division of Neurology, Department of Medicine, University of Alberta; Edmonton, Alberta, Canada, 8 Microsoft Research; Redmond, Washington, United States of America, 9 The Faculty of Data and Decision Sciences, Technion; Haifa, Israel, 10 Department of Applied Mathematics and Theoretical Physics, University of Cambridge; Cambridge, United Kingdom, 11 The Alan Turing Institute; London, United Kingdom, 12 Independent Researcher; United States of America

These authors contributed equally to this work.
\* <u>Imccoy@ualberta.ca</u>

# Abstract

Despite significant technical advances in machine learning (ML) over the past several years, the tangible impact of this technology in healthcare has been limited. This is due not only to the particular complexities of healthcare, but also due to structural issues in the machine learning for healthcare (MLHC) community which broadly reward technical novelty over tangible, equitable impact. We structure our work as a healthcare-focused echo of the 2012 paper "Machine Learning that Matters", which highlighted such structural issues in the ML community at large, and offered a series of clearly defined "Impact Challenges" to which the field should orient itself. Drawing on the expertise of a diverse and international group of authors, we engage in a narrative review and examine issues in the research background environment, training processes, evaluation metrics, and deployment protocols which act to limit the real-world applicability of MLHC. Broadly, we seek to distinguish between machine learning ON healthcare data and machine learning FOR healthcare—the former of which sees healthcare as merely a source of interesting technical challenges, and the latter of which regards ML as a tool in service of meeting tangible clinical needs. We offer specific recommendations for a series of stakeholders in the field, from ML researchers and clinicians, to the institutions in which they work, and the governments which regulate their data access.

B. is an employee of IBM Research. J.G. is a 2022 Robert Wood Johnson Foundation Harold Amos Medical Faculty Development Program and declares support from RSNA Health Disparities grant (#EIHD2204), Lacuna Fund (#67), Gordon and Betty Moore Foundation, and NIH (NIBIB) MIDRC grant under contracts 75N92020C00008 and 75N92020C00021. U.S. is partially funded by the Israeli Science Foundation grant no. 1950/19. L.A.C. is funded by the National Institute of Health through NIBIB R01 EB017205. T.N. is an employee of Microsoft Research. M.vdS. is supported by the Office of Naval Research (ONR). The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

**Competing interests:** L.A.C. is the Editor-in Chief of PLOS Digital Health.

### Author summary

The field of machine learning has made significant technical advancements over the past several years, but the impact of this technology on healthcare practice has remained limited. We identify issues in the structure of the field of machine learning for healthcare which incentivise work that is scientifically novel over work that ultimately impacts patients. Among others, these issues include a lack of diversity in available data, an emphasis on targets which are easy to measure but may not be clinically important, and limited funding for work focused on deployment. We offer a series of suggestions about how best to address these issues, and advocate for a distinction to be made between "machine research performed ON healthcare data" and true "machine FOR healthcare". The latter, we argue, requires starting from the very beginning with a focus on the impact that a model will have on patients. We conclude with discussion of "impact challenges"— specific and measurable goals with an emphasis upon health equity and broad community impact—as examples of the types of goals the field should strive toward.

### Introduction

The 2012 paper "Machine Learning that Matters" [1] introduced Impact Challenges as a mechanism for opening discussion in the machine learning (ML) community about the limitations in datasets, metrics, and the impact of results in the context of their originating domains. The article presents the ML community as overly concerned with theoretical, benchmark-focused work detached from real-world problems, and ultimately failing to address tangible issues. Despite the incredible progress that has been made in ML [2], particularly in the field of large generative models [3], many of the paper's original criticisms continue to ring true. We contend that these issues are of particular concern in the field of machine learning for healthcare (MLHC).

Despite its challenges, such as the safety-critical [4] nature of decisions and the complex multi-stakeholder [5] environments, healthcare is in many ways an ideal setting for the application of machine learning [6]. The amount of data involved is vast [7], and the setting is deeply meaningful. Research in this space can have immense value [6]–from providing timely, more precise and more objective decision support for providers and patients [8,9], to cognitive offloading for overburdened healthcare workers [10,11], to improving health system efficiency and population health outcomes [12].

Yet, in our opinion, there exists a stark contrast between the stated ambition (and associated hype) of the MLHC field and its limited degree of meaningful impact to date. The approach of the field to the COVID-19 pandemic serves as an illustrative example. With the pandemic affecting countries world-wide, there were multiple coordinated efforts to collect and share data for ML-based COVID-19 diagnosis [13]. However, despite the generation of several hundred published predictive models, systematic reviews found severe roadblocks to the use of these models in clinical practice [14,15]. Particularly, only a handful were promising enough for deployment in real clinical settings [16].

We believe that this example, while an extreme case, is not an outlier. While technical tools in the space of ML for healthcare have made great leaps [17], social aspects of such tools in practice-be it their design, development or deployment-are often afterthoughts [18–20]. Models are too often developed without appropriate scrutiny of underlying data and subsequent outputs [14,21]. Therefore, machine learning models learn and ultimately reproduce the biases of society [22] at large, as reflected in racial differences in care patterns [23–27], poor



#### Fig 1. Machine learning for healthcare impact framework and recommendations.

https://doi.org/10.1371/journal.pdig.0000474.g001

performance of clinical devices such as oxygen saturation probes for patients with dark skin [28,29], and numerous other aspects of the broader care system [30].

The MLHC field has, in our opinion, failed to address these issues, and has indeed exacerbated them through structures which broadly reward technically novel innovations over tangible clinical impacts. Throughout this paper, we will explore issues in the background environment, development processes, research evaluation, and clinical deployment of MLHC projects. We seek to critically analyze misaligned incentives which have led to a stark disconnect between our field's research and its broader societal impact. While several prior works [6,31–37] have addressed this topic to varying degrees, we intend this work to summarize gaps and pain points, and we present recommendations to re-center the focus of machine learning as applied to healthcare.

We conclude with a series of renewed impact challenges, hoping to re-center the notion that the core purpose of MLHC is to achieve equitable impact on meaningful clinical out-comes. All aspects of MLHC—background environment, development processes, evaluation metrics, and deployment efforts-must be understood in terms of how they advance or hinder such tangible goals. An overview of our suggestions can be seen in Fig 1.

### Environment: Background for building

The context in which research is conducted serves as the foundational platform, influencing the formation, development, and impact of research findings. Particularly in the rapidly-evolving field of MLHC, the broader research environment sets the stage for how and what kind of knowledge is generated. It is crucial to recognize that the nature and quality of data, the accessibility of this data, the inherent biases present, and the diversity of those conducting research all converge to form this dynamic milieu. The importance of this research background extends beyond the realm of technical advancements, shaping the ethical, societal, and practical implications of AI technologies.

### Understanding social determinants and data biases

Disparities in health and healthcare are present along numerous axes, including but far from limited to race, gender, socioeconomic status, and nationality [27]. Uncritically trained MLHC models tend to learn, reproduce, and further entrench these disparities—as shown in the prominent case of an algorithm which used past spending received as a proxy for health needs and further disadvantaged Black patients [27]. It is not enough to simply remove these attributes from the datasets, as ML models have demonstrated a concerning degree of ability to ascertain protected characteristics even in the absence of obvious proxies [38]. The data produced by social processes cannot be understood in isolation from those processes [39]. Similarly, work must be done not merely to eliminate the biases in health data, but to actively characterize and counteract the conditions which generate these biases in the first place. Researchers must seek to achieve "data empathy" [40], and understand the intensely human processes and stories that data—particularly data in healthcare—embed [41].

We recommend:

- Work to actively characterize and counteract underlying disparities and social determinants of health embedded in healthcare data, particularly in international contexts.
- Engage critically with processes underlying data production, and seek both upstream and downstream measures to ensure that all patients are equitably represented and impacted.
- Ensure that patient perspectives and insights are represented during data creation, validation, and release phases.
- Pursue the goal of "data empathy", ensuring that the human stories underlying medical data are not forgotten amidst the abstracting processes of MLHC.

#### Improving diversity among researchers

Illness impacts us all, but the burdens of both disease and substandard care tend to fall most heavily upon those who are already marginalized [42]. However, neither the demography nor the processes of the MLHC field reflect this reality. Both the broader MLHC field at large and the subfield of fairness research are disproportionately composed of male, white, and Asian researchers from a small set of high-income countries [43]. The research community as a whole also skews both younger and more able bodied than broader patient populations [44]. The result is a field with significant blind spots, and a broader failure to adequately consider and align with the voices of those who have greatest need [45]. In order to truly achieve impact, the MLHC field must embrace diversity in its deepest sense, with researchers from a wide range of racial, gender, and national backgrounds.

The need for diversity also includes the training background and professional roles of those involved, and a wide range of stakeholders must be actively invited in at every stage of the research process. In this regard, the MLHC field should engage with and build upon existing "patient engagement" practices in clinical trials [46], which seek to ensure that research is aligned with patient needs [47–49]. With MLHC envisioned to impact a broad range of health-care contexts, all of those involved may have valuable insights in co-designing projects [50]—including patients, caregivers, and practitioners historically underrepresented in the academic research process such as homecare workers or nursing aides [51].

We recommend:

• Promote MLHC training to a wide range of health and computer science trainees, with a particular emphasis upon members of underrepresented groups.

- Carry out international events such as datathons [52] to build global collaborative networks, ensuring that the agenda of the field is not controlled by those in a narrow set of high-income countries.
- Actively invite patient and caregiver stakeholders into all levels of the process, ensuring that procedures and outputs are aligned with tangible patient needs.
- Emphasize diversity in training and practice contexts (such as inclusion of clinicians from minority-serving institutions in high income countries), in addition to diversity in background.
- Improve the upstream training pipeline through investment in developing foundational data science capacity at minority-serving institutions such as historically Black colleges and universities,

### Fostering diverse dataset creation

Machine learning thrives on data, yet this is not fully reflected in current conference proceedings, where papers that discuss the development of datasets are rare. Some venues have emerged (such as the NeurIPS Datasets and Benchmarks track) [53], but they remain relatively novel and rare. The causes of this are multifactorial. Developing datasets requires significant effort, particularly in complex domains such as healthcare [54], and requires collaborations [55], resources, and funding which are often limited to only the largest of institutions [56]. Concerns regarding patient privacy and downstream data use further limit these collaborations specifically in the healthcare context.

As a consequence, a significant proportion of machine learning work tends to be excessively focused on a few publicly available datasets [57]. When these datasets are not sufficient for data hungry models, researchers can combine multiple datasets resulting in the so-called Frankenstein datasets [14], which are difficult to audit and can have misleading assumptions when a class label is assigned. ML works related to healthcare are subsequently limited in scope to a small set of problems such as prediction of mortality [58] or hospital length of stay [59], while widespread issues of critical importance such as maternal mortality [60,61] or discrepancies in healthcare access remain underexplored [62]. Further, these few datasets are skewed toward representing the subset of patients served by large institutions in affluent geographic locations [63]. Subsequently, algorithms developed using these datasets may serve to further exacerbate disparities to the disadvantage of underrepresented patient populations.

There are a growing range of initiatives designed to increase the diversity of dataset creation. NIH projects such as AIM-AHEAD [64] and BRIDGE2AI [65] aim to expand health data science capacity throughout the United States. Capacity in developing countries is being fostered by programs such as the NIH DS-I Africa program [66], as well as initiatives from the Gates Foundation [67] and Wellcome Trust [68]. It is critical to ensure that these capacity building efforts are not merely limited to the provision of funding, but also include work to foster a data sharing culture, address local privacy concerns [69,70], and promote multidisciplinary collaboration [71]. Local leadership and agency in this regard must remain paramount, in order to avoid perpetuation of global health power imbalances. In addition, datasets from underrepresented sources must be actively included in the mainstream of MLHC research, so that cutting-edge core models and methodologies are representative of *all* patients.

We recommend:

• Recognize the critical importance of dataset creation work in funding, hiring, and promotion contexts.

- Focus the creation of novel datasets on rectifying existing issues of patient underrepresentation.
- Provide detailed dataset descriptions (such as those described in "Datasheets for Datasets" [72,73]) that inform data harmonization practices, especially when the data use is different from its original intended use.
- Ensure that datasets created in low resource settings are included in the mainstream advancement of MLHC, rather than being limited to regional use.

### Opening data and increasing accessibility

Where robust and comprehensive datasets do exist, they are often subject to significant restrictions which limit their accessibility and usefulness to the MLHC community [57]. This may be due to concerns from host health organizations regarding patient privacy or re-identification risk [74,75]. There may also be a desire on the part of dataset creators to maintain exclusivity and reap the professional rewards of their hard work in creating the dataset in the first place.

Experience with existing open datasets, such as the Medical Information Mart for Intensive Care (MIMIC) [76], has demonstrated both sets of concerns to be often overblown [77]. With respect to privacy, de-identification measures are highly effective in practical terms, and there have been no known instances of re-identification of individuals in this dataset since its initial release in 1996 [78]. With respect to research output, experience has shown that open datasets beget a synergistic momentum, with multiple groups able to both collaborate and compete to drive the field forward. As the 5500+ citations of the MIMIC-III database have demonstrated [76], there is no shortage of clinical questions to be answered on a given set of data.

We recommend:

- Promote the creation of FAIR (Findable, Accessible, Interoperable, and Reusable) [79] datasets in healthcare, alongside the sharing of relevant supportive resources and code for dataset cleaning and optimization.
- Expand privacy-preserving efforts in areas such as vectorization, synthetic data, and federated learning, while remaining wary of their limitations and favoring completely open data sharing where possible.
- Ensure that efforts toward open data are inclusive of a broad range of healthcare settings, both internationally and within a given country. This includes not only large academic tertiary hospitals, but also regional hospitals, and alternative care settings such as community clinics.

### Building foundational tools for MLHC success

If the benefits of machine learning in healthcare are to be realized equitably, they must be realized at scale within numerous diverse clinical contexts. Where this localization does occur, it is often performed in labor-intensive ways by high-cost teams of experts, who often find themselves reinventing the proverbial wheel. For scalable success, MLHC must embrace the spirit of the open source movement, and work collaboratively to develop open automated methods to assist in model development, localization, and validation [80]. A notable and successful effort in this direction is the work done by the OHDSI consortium [81]. The field must recognize and reward such infrastructural work, given its critical importance to achieving scalable impact.

- Develop and standardize ML methods for data harmonization, reducing the significant existing barriers to bringing together data from multiple clinical sources.
- Require sharing of data cleaning and preprocessing code, in addition to final model development code, with detailed performance breakdowns for data subgroups using reporting tools like model cards [82].
- Create automated systems for comparing methods and facilitating evaluation.
- Create open-sourced autoML frameworks to automatically compare developed models systematically on important metrics. [80]
- Contribute to developing robust open frameworks for enduring research data sharing and distribution (such as dataverse). [80]

### **Process: Training for impact**

In the ever-evolving landscape of machine learning in healthcare (MLHC), the importance of process—how we frame, conduct, and evaluate our research—is paramount. It's not merely a question of pursuing advanced technology as abstractly defined. Rather the process itself holds the key to ensuring the relevance, applicability, and ethical integrity of our findings. The forth-coming section explores this fundamental concept, illustrating how meaningful task selection, ambitious benchmarking, consistent responsibility standards, and operationalized fairness are critical in shaping the impact of MLHC research. By anchoring our efforts in these principles, we transition from myopic technical novelty to a more balanced pursuit of solutions that are patient-centered, equitable, and ultimately transformative for healthcare.

#### Selecting meaningful tasks

Given the field's hunger for technical novelty, MLHC research can sometimes focus on "solutions in search of problems" [83]. Researchers may regard healthcare primarily as a source of complex interesting data with compelling real-world justifications. There can be a tendency to focus on implementing the most interesting and novel techniques on healthcare data, rather than solving the most urgent and tangible healthcare problems. While "machine learning on healthcare data" can be a meaningful way to drive forward ML research, it must not be confused with true "machine learning for healthcare". Problems in healthcare are generally complex and context-specific.

We recommend:

- Distinguish between "machine learning on healthcare data" and "machine learning for healthcare problems", recognizing that the technical novelty of the former does not innately translate to impact on the latter.
- Engage with clinical stakeholders early in the clinical process, and working from problem to solution with earnest evaluation of whether a particular method—or machine learning at all —is most appropriate to address the problem.

#### Moving past the status quo

A common approach in MLHC is using clinical data to predict variables corresponding with a clinician's decision, such as the diagnostic label of an X-ray [84,85], or the decision to prescribe a medication [86]. Such approaches implicitly treat that clinical assessment as the ground truth, and in so doing establish contemporary clinician-level performance as a ceiling. This is

problematic given the well-documented fallibility of human clinicians [87], and the pervasive biases which further worsen the quality of care for underserved patients [88,89].

While learning from clinicians (particularly ensembles of clinicians) is important, MLHC should strive where possible to surpass the quality and accuracy of existing clinical practice. A particularly illustrative example was provided by Pierson et al. [90], who demonstrated that training a knee X-ray algorithm to predict *patient-reported pain* rather than *radiologist-assessed arthritis burden* produced a model which eliminated pre-existing racial disparities in X-ray assessment.

We recommend:

- Focus wherever possible on prediction of either objective ground truth metrics such as mortality or relevant patient centered metrics such as reported pain, rather than strict agreement with human clinician assessments alone.
- Engage in ambitious attempts to elevate existing clinical standards, rather than simply striving to meet the baseline status quo.

#### Establishing consistent responsibility standards

Current MLHC development lifecycles consider "performance improvement" in a given core metric such as area under the precision-recall curve (AUPRC) or accuracy metrics as the primary target [91–93]. The field is focused upon defining and carrying out narrow tasks, with the broader context and ultimate impact of the model often considered an afterthought [94]. This reality shapes every step of the MLHC process, from data collection, to model development, to validation—with issues of responsible AI such as fairness, accountability, and transparency considered after the fact if at all. While significant literature explores standards related to these issues [72,95–97], they are implemented in a patchy and inconsistent manner which renders comparison and accountability between institutions difficult.

We recommend:

- Establish and propagate industry standards for responsible AI development, incorporating patient-centric and ethical values. [83]
- Dethrone narrow performance metrics as the primary assessment method, and consider them alongside clinical impact and responsible AI issues throughout the development process.

#### **Operationalizing fairness**

Fairness and equity are often posited as goals of MLHC development, however the terms are often used vaguely and without clear meaning, as previous criticism has noted [98]. Failure on the part of both the MLHC community and the broader regulatory apparatus to develop a clear set of expectations in this regard enables research with a wide range of actual impact to be passed off as sufficiently "fair". Far too often, the equity-focused analyses performed are only cursory, hidden away in supplemental tables and hardly engaged within the main body of a paper. Regardless of the specific consensus definition of the field, if fairness is to be achieved in MLHC, it must be baked into the process rather than painted on after the fact. It must be understood in concrete terms, and have the power to meaningfully shape the development process.

- Pursue consensus definitions of fairness in MLHC contexts, both procedural as well as outcome-based.
- Engage in subgroup analysis of the outputs of all MLHC projects, in order to understand and engage in dialogue and remediation regarding relevant disparities.
- Develop and hold projects to explicit standards regarding fairness, equity, and maximally permissible performance disparities between demographic groups.
- Render fairness constraints an integral part of training processes, rather than an ad-hoc or after-the-fact correction.

### **Evaluation: Re-aligning incentives**

Recognizing that innovation isn't merely about technical novelty but also about potential clinical impact can lead to a paradigm shift in how research is conducted and valued. The lure of common metrics of success can lead researchers to overlook the complex realities of realworld clinical contexts, and the sometimes subtle but significant disparities that exist between controlled research environments and the day-to-day chaos of healthcare provision. Whether through completing moonshot projects or solving overlooked last-mile challenges, MLHC researchers must work to create health equity, rather than merely managing bias. We advocate for a broadened perspective on the value of innovation, a more nuanced understanding of real-world applicability, and an appreciation for audacious endeavors, thus forging a path towards a more impactful MLHC landscape.

#### Recognizing advances for more than just algorithmic/ technical novelty

Innovation is often equated with novelty in academic research, particularly in the increasingly intense competition for conference reviewer attention and approval. Commending research primarily for its originality encourages new and creative methods which may be ultimately only incremental in impact. At the same time, this philosophy acts to relatively disincentivize the unglamorous work necessary to develop an idea fully and address all of the various challenges which arise in its deployment in the clinical context. The result is an excess of incomplete research agendas, as researchers face rapidly diminishing returns for the additional follow-up work required to fully develop the implications of an initially novel work.

This issue is not merely in the realm of academic recognition, as both governmental grants and industry funding tend to follow a similar pattern. Given the increasing costs of model training, assessments of what will be fundable act to shape and ultimately bias research in directions which may be more attention-grabbing than impactful. Further, projects which are funded for technical novelty often lack access to the necessary follow-on funding to be maintained through the full life cycle and achieve impact.

- Recognize that novelty has multiple aspects by creating explicit research tracks highlighting technical significance and clinical relevance.
- Provide opportunities for follow-on funding to carry ideas forward to implementation and tangible impact.
- Conduct retrospective analyses of trends in machine learning applications for health [99], and introspect on areas that might require more research focus.

### Moving on from clinically unimportant metrics

MLHC research tends to focus on label prediction accuracy metrics such as sensitivity, specificity, and the precision-recall curve [91–93]. However, accuracy when predicting labels on a curated dataset does not always translate to accuracy in real-world clinical settings when deployed, often due to dataset shift [100,101] caused by critical and inescapable differences between the populations used in testing versus deployment [102]. More substantially, all accuracy metrics should be considered with respect to a clinical goal and a clinical workflow. Even an accurate prediction may have little clinical value [103] in certain circumstances—it may be too late, too early, about an event that cannot be prevented, or add little useful information to existing clinical assessments. Thus, the value of accuracy metrics as proxies for clinical relevance is highly variable and ultimately task- and context-specific.

We recommend:

- Develop and validate metrics in line with clinical goals and workflows, with an awareness of when information will augment meaningfully what clinicians already know [104].
- Validate clinical targets and metrics via clinical trials focused on tangible patient-relevant endpoints such as mortality or disability reduction.
- Pursue causal modeling for automated assessment of whether a given alert is likely to impact clinical decision-making.
- Evaluate machine learning solutions against already existing methods used in relevant healthcare settings as well as simple non-machine learning baselines.
- Monitor performance after deployment to identify the presence of dataset shift and help inform decisions about when model re-training is needed.
- Evaluate models in the local context of their intended deployment, recognizing that performance can be sensitive to subtle specifics.

### **Encouraging moonshots**

Stepping outside the comfortable bounds of topping established benchmarks purely *in silico* exposes a project to a wide range of uncertainties, and the most potentially impactful MLHC projects are often also the riskiest. They require seeking the approval of a wide range of stake-holders, and they often play out across extended multi-year timelines [105]. This can conflict with the career incentives of MLHC academics, particularly those on short tenure timelines, as well as students with brief PhD or even briefer MSc timelines. Researchers may fear—and with some justification—that pursuing such projects places the eggs of their career into a single fragile basket. If the field is going to succeed in having impact, this dynamic must be reversed. We recommend:

- Recognize the insights gained from ambitious projects, even (or especially) if they ultimately are not successful.
- Encourage publication of process descriptions and intermediate results [106] for prolonged, multi-stage development and implementation.
- Allocate grant funding to ambitious projects with longer timelines, aiming at real-world clinical benchmarks and equitable outcomes.
- Publish "lessons learned" from projects that may not have succeeded at their original goals but nevertheless have insights of value for the rest of the field.

### Deployment: From code to clinic

The path from theory to practice is paved with often-overlooked aspects, including rigorous engineering, a profound understanding of human processes, acknowledgment of system limitations, anticipation of ongoing impact, and a commitment to parallel valuation of validation and innovation [105]. Each of these dimensions bears a unique set of considerations and requires explicit focus to ensure successful, safe, and sustainable implementation of MLHC. In the subsequent section, we delve into these complexities and offer recommendations that aim to foster an approach that is not just technically sound, but also holistic, human-centered, transparent, and future-oriented. Through these concerted efforts, we aspire to enhance the reliability, efficacy, and ultimately, the patient outcomes of MLHC deployment in diverse healthcare environments.

### Valuing engineering effort

Machine learning models require significant work to be deployed in practice, especially in high-stakes domains such as healthcare [107,108]. The scientific experiments accompanying the proposal of a new model, while necessary, are not sufficient to deem a model useful and practical [14]. Deployment involves significant work [107,108], such as cross-contextual validation, development of platforms or APIs for model access, online model monitoring, and data collection for further model improvements. All this accompanying work may, unfortunately, not be recognized as part of the research process, despite the fact that it is mandatory to achieve wide model use. This work is often separated out as "engineering" and deemed uninteresting in the academic context despite its presence as an essential component of any translation effort. This is particularly problematic in healthcare, given the variability and complexity of both the digital systems and sociotechnical structures involved.

The MLHC community should learn from and build upon preceding efforts in public health and health promotion in this regard. The Reach, Effectiveness, Adoption, Implementation, and Maintenance (RE-AIM) framework [109] is one robust example, which seeks to find a balance between the internal and external validity of projects by highlighting these challenges from the outset of a project. In particular, the MLHC community must come to regard the effort necessary to adapt to existing, complex healthcare contexts as core to, rather than ancillary to, any new project.

We recommend:

- Develop research tracks encouraging "last-mile" work, with an emphasis on deployment and impact evaluation.
- Create specific conference venues dedicated to engineering problems in machine learning for healthcare.
- Seek active collaboration with colleagues with expertise in data engineering, software engineering, systems engineering, bioinformatics, and related subfields.
- Develop and adopt standards that reduce the technical burden of organizations to deploy and monitor algorithmic performance in the real world setting.
- Utilize frameworks such as RE-AIM [109] to conceptualize and prepare for the work which must be done to adapt MLHC initiatives to real-world contexts.

### **Considering human processes**

While many MLHC projects are designed and reported in controlled computational isolation, all clinical processes involve the variability and complexity inherent to human involvement

[110]. Numerous questions arise at this juncture. How will model outputs be understood by clinicians? What does it take for a model's output to be trusted? Will the presence of an inaccurate model recommendation cause clinicians to second-guess their better judgment [111]? The MLHC field must work to further develop the multidisciplinary work that has been done to characterize human-machine systems, and the habits that clinicians build when working with machines. Well-characterized issues include automation bias [112–114] (overreliance on machine recommendations), algorithm aversion [115,116] (underreliance on machine recommendations), and alarm fatigue [117] (becoming overwhelmed by the frequency of alerts). Any approach which understands an ML model in isolation is fundamentally incomplete, and models developed in such isolation will fail to have tangible clinical impact. We recommend:

- Work to understand the role of algorithms amidst the broader human-machine system [118].
- Engage clinicians, nurses, patients, and other end-users of systems in order to understand how model outputs are understood, and how their presentation can be optimized.
- Collaborate with colleagues in fields such as medical anthropology, human factors engineering, psychology, sociology, feminist techno-science, and user experience design.
- Establish processes for ongoing monitoring of clinician feedback, ensuring that the practical usability of models is continuously optimized.
- Ensure that appropriate backup procedures are in place to recognize and respond to model downtime or performance deterioration.
- Continue training clinicians for model-free circumstances, in order to avoid deskilling [119].

### Identifying system limitations

Current MLHC research is biased toward demonstration of the unique and novel capabilities of a model at its best. Yet, of equal importance is a robust understanding of what a given model *cannot* do. Characterizing and elaborating upon the particular failure cases of a model is essential to safe and impactful deployment, and enables amelioration measures such as learning when to defer to human clinician assessments [120,121]. Identification of limitations is also essential to ensure that a model is not uncritically deployed outside of the initial scope for which it was developed and validated.

- Reward researchers in peer review and publication processes for honest and thoughtful characterization of the limitations of a given model architecture.
- Ensure that regulatory processes require clear establishment of model limitations, and that these are consistently made clear to health systems and model end users.
- Work to generate standardized adversarial assessments [122] to probe the vulnerabilities of a given clinical model.
- Utilize standardized model reporting techniques, such as "Model Cards" [82] in order to ensure that model training contexts, targeted uses, and limitations are clearly disclosed.

### Building for continued impact

Rewards in academic MLHC research are heavily skewed toward the initial design and deployment of a model [100], whereas the ultimate clinical impact accumulates gradually over an extended period of time. The continued maintenance and execution of an MLHC project is a complicated endeavor in its own right, with issues such as dataset shift impacting model accuracy, and advancements in treatment protocols possibly impacting a project's overall appropriateness. Anticipating and addressing such factors must be considered an integral part of the MLHC enterprise, rather than being relegated as an unimportant ongoing maintenance task for the end-user [123]. Projects which are initiated but not maintained may even act as a net harm to patients overall, as processes may be altered and resources may be re-allocated in ways that are not easily reversed.

We recommend:

- Encourage the publication and dissemination of longitudinal assessments reporting retrospectively on the impact of models years after initial deployment.
- Establish guidelines and checklists to update and/or retrain models [123] in response to changing data and process environments.
- Establish institutional monitoring teams and processes for the longitudinal assessment and adjustment of clinical models for issues such as dataset shift.
- Offer rewards for long-lived, successful deployments of MLHC projects, as well as terminating projects assessed to have harms or negative impacts.

#### Valuing generalization on par with innovation

Even where MLHC projects *are* successfully implemented at a single site, machine learning models experience a notorious degree of difficulty generalizing outside of their initial contexts. If MLHC is going to have an impact on patients broadly, there is significant work to be done in ensuring that the proverbial wheel does not require re-inventing in every hospital and clinic around the world [124]. At the other end of the spectrum, some models have been widely deployed without any significant effort toward validation, with deleterious consequences [124]. Yet as has been described with the replication crisis in psychology and other sciences, academic research conveys relatively little reward for the co-equal work necessary for validation.

Further, validation remains often project and context-specific, with a paucity of generalizable methods for comparison between projects. When considered in terms of patient impact, however, performance at each additional site is equally important to performance at the index institution. Methods such as "realist evaluation" [125] are highly useful in this regard, providing for formal structures to evaluate "what works, for whom, and in what circumstances" [126]. The framework ensures that contextual factors (such as the individuals involved, and their broader infrastructure) and context-specific mechanisms are explicit targets of evaluation and assessment.

- Dedicate increased focus to the techniques of transfer learning and local fine-tuning when necessary.
- Develop consistent interoperability standards, and methods for understanding the shifts in model performance as adapted to different contacts.

- Encourage publication of validation studies, with particular description of pitfalls faced in local translation.
- Create pipelines for the systematic comparison and validation of methods [80].

### Impact challenges in machine learning for healthcare

We believe that MLHC must be oriented first and foremost toward its impact upon the health of individual patients and the community at large. In the same vein as Wagstaff (2012) [1], we seek to offer a series of impact challenges which highlight tangible, meaningful goals for progress in the field [127].

- A model deployed in an American urban hospital recognizes high-risk prenatal patients, recommends interventions, and successfully reduces the Black-white maternal mortality disparity by > = 50%.
- 2. A model deployed and maintained at a single center to predict patient deterioration [128] continues to maintain > = 95% of its initial predictive performance and impact on mortality across all demographic groups over a 10-year timeframe.
- 3. Data pipelines and transfer learning methods are used to deploy a chest x-ray diagnostic model [129] with significant clinical impact across urban and rural hospitals in three countries in Sub-Saharan Africa.
- 4. A "second opinion" diagnostic language model [130] for common clinical ailments is actively consulted as a part of regular outpatient clinical workflow with a >90% physician satisfaction rate with answer usefulness across multiple health systems in diverse socioeconomic contexts.

The purpose of these challenges is not to be prescriptive, but rather to highlight examples of concrete end-goals which the broader MLHC enterprise must be in service of. The structural recommendations highlighted elsewhere in this paper aim to create the contexts and processes —the "where", "who" and "how"—necessary to work toward such end-goals. Crucially, these goals do not need to be achieved from beginning to end by a single group, and MLHC endeavors do not need to be limited to grand projects. Rather, they are meant to frame and guide the field as a whole—that is, even smaller projects should be conceived, understood, and implemented in terms of how they move towards ambitious goals with tangible clinical impact.

Seeking to achieve (1), for example, would require moving beyond the task of simply predicting patient outcomes, and toward building MLHC as an integral part of a broader complex system considering the social dimensions of health. It would require not only technical expertise, but also consideration of patient needs and clinical barriers in a specific socio-technical context. In doing so, it would necessitate the collaborative involvement of members of a racially, culturally, and disciplinarily diverse team, which requires the expansion of inclusive training pathways, and efforts to promote MLHC among those interested in maternal health. Further, it would require the upstream expansion of data pathways to gather data outside of traditional academic centers, and the creation of open datasets necessary for fundamental ML methods to be applied to the field of maternal health.

Both machine learning and healthcare are highly complicated topics in isolation, and MLHC is in many ways even more so. We believe that a healthy and thriving MLHC community and research apparatus is one that is able to consistently, aim toward and ultimately achieve goals such as those embodied by these challenges. It is incumbent upon all of us

working in this field to work toward building such an apparatus—working not only to demonstrate technical novelty on data gathered amidst the background of healthcare, but to consistently generate solutions which work for all.

### Conclusion

Despite major advancements in the underlying technology, we contend that the impact of MLHC on meaningful clinical problems has been limited. This limitation in part reflects the inherent difficulty of deploying ML solutions in such a high-complexity and high-stakes domain. However, it also reflects significant issues in the structure and culture of the field of MLHC at large.

Data access [107,131] is woefully limited and inadequate to reflect the diversity of patient populations [132]. The research process often fails to select and engage with meaningful tasks. Evaluation tends to prize novelty over impact [1], and the field gravitates toward simplistic metrics. A majority of research and development efforts focus on technical "novel" model development rather than data collection and deployment. Underpinning this all, significant issues of bias in both data and process lead to the production of models which may only exacerbate existing healthcare disparities [27].

As we have discussed throughout this paper, these problems are real and substantial, but far from intractable. We are hopeful that this article may help to spark reflection and conversation within the MLHC field. We must work to distinguish between "machine learning research performed on healthcare data" and "machine learning for healthcare". We must re-orient ourselves toward the grand challenges which exist, and the tangible steps required to address them. We must strive to be a field defined first and foremost by our impact on the lives of patients and their families—that is, to engage in machine learning for healthcare that matters.

### **Author Contributions**

Conceptualization: Ioana Baldini, Leo Anthony Celi, Tristan Naumann, Uri Shalit.

- **Investigation:** Aparna Balagopalan, Ioana Baldini, Leo Anthony Celi, Judy Gichoya, Liam G. McCoy, Tristan Naumann, Uri Shalit, Mihaela van der Schaar, Kiri L. Wagstaff.
- Methodology: Aparna Balagopalan, Ioana Baldini, Leo Anthony Celi, Judy Gichoya, Liam G. McCoy, Tristan Naumann, Uri Shalit, Kiri L. Wagstaff.
- Supervision: Ioana Baldini, Leo Anthony Celi, Judy Gichoya, Tristan Naumann, Uri Shalit, Mihaela van der Schaar, Kiri L. Wagstaff.
- Visualization: Aparna Balagopalan.
- Writing original draft: Aparna Balagopalan, Ioana Baldini, Leo Anthony Celi, Judy Gichoya, Liam G. McCoy, Tristan Naumann, Uri Shalit, Mihaela van der Schaar, Kiri L. Wagstaff.
- Writing review & editing: Aparna Balagopalan, Ioana Baldini, Leo Anthony Celi, Judy Gichoya, Liam G. McCoy, Tristan Naumann, Uri Shalit, Mihaela van der Schaar, Kiri L. Wagstaff.

#### References

- 1. Wagstaff K. Machine Learning that Matters In Proceedings of the 29th International Conference on International Conference on Machine Learning (pp. 1851–1856).
- 2. Radford A, Kim JW, Hallacy C, Ramesh A, Goh G, Agarwal S, et al. Learning Transferable Visual Models From Natural Language Supervision. In: Meila M, Zhang T, editors. Proceedings of the 38th

International Conference on Machine Learning. PMLR; 18–24 Jul 2021. p. 8748–63. (Proceedings of Machine Learning Research; vol. 139).

- Biswas SS. Role of Chat GPT in Public Health. Ann Biomed Eng. 2023 May; 51(5):868–9. <a href="https://doi.org/10.1007/s10439-023-03172-7">https://doi.org/10.1007/s10439-023-03172-7</a> PMID: 36920578
- Killian T. W., Parbhoo S. & Ghassemi M. Risk Sensitive Dead-end Identification in Safety-Critical Offline Reinforcement Learning. Transactions on Machine Learning Research (2023).
- Schiavone F, Mancini D, Leone D, Lavorato D. Digital business models and ridesharing for value cocreation in healthcare: A multi-stakeholder ecosystem analysis. Technol Forecast Soc Change. 2021 May 1; 166:120647.
- Ghassemi M, Naumann T, Schulam P, Beam AL, Chen IY, Ranganath R. A Review of Challenges and Opportunities in Machine Learning for Health. AMIA Jt Summits Transl Sci Proc. 2020 May 30; 2020:191–200. https://doi.org/10.1001/jama.2017.18391 PMID: 32477638
- Baro E, Degoul S, Beuscart R, Chazard E. Toward a Literature-Driven Definition of Big Data in Healthcare. Biomed Res Int. 2015 Jun 2; 2015:639021. <u>https://doi.org/10.1155/2015/639021</u> PMID: 26137488
- O'Connor PJ, Sperl-Hillen JM, Rush WA, Johnson PE, Amundson GH, Asche SE, et al. Impact of electronic health record clinical decision support on diabetes care: a randomized trial. Ann Fam Med. 2011 Jan-Feb; 9(1):12–21. https://doi.org/10.1370/afm.1196 PMID: 21242556
- Levin S, Toerper M, Hamrock E, Hinson JS, Barnes S, Gardner H, et al. Machine-Learning-Based Electronic Triage More Accurately Differentiates Patients With Respect to Clinical Outcomes Compared With the Emergency Severity Index. Ann Emerg Med. 2018 May; 71(5):565–74.e2.
- Tuli S, Basumatary N, Gill SS, Kahani M, Arya RC, Wander GS, et al. HealthFog: An ensemble deep learning based Smart Healthcare System for Automatic Diagnosis of Heart Diseases in integrated IoT and fog computing environments. Future Gener Comput Syst. 2020 Mar 1; 104:187–200.
- Thakkar D, Ismail A, Kumar P, Hanna A, Sambasivan N, Kumar N. When is Machine Learning Data Good?: Valuing in Public Health Datafication. In: Proceedings of the 2022 CHI Conference on Human Factors in Computing Systems. New York, NY, USA: Association for Computing Machinery; 2022. p. 1–16. (CHI '22).
- 12. Ehrmann DE, Gallant SN, Nagaraj S, Goodfellow SD, Eytan D, Goldenberg A, et al. Evaluating and reducing cognitive load should be a priority for machine learning in healthcare. Nat Med. 2022 Jul; 28 (7):1331–3. https://doi.org/10.1038/s41591-022-01833-z PMID: 35641825
- Shuja J, Alanazi E, Alasmary W, Alashaikh A. COVID-19 open source data sets: a comprehensive survey. Appl Intell (Dordr). 2021; 51(3):1296–325. <u>https://doi.org/10.1007/s10489-020-01862-6</u> PMID: 34764552
- Roberts M, Driggs D, Thorpe M, Gilbey J, Yeung M, Ursprung S, et al. Common pitfalls and recommendations for using machine learning to detect and prognosticate for COVID-19 using chest radiographs and CT scans. Nature Machine Intelligence. 2021 Mar 15; 3(3):199–217.
- Wynants L, Van Calster B, Collins GS, Riley RD, Heinze G, Schuit E, et al. Prediction models for diagnosis and prognosis of covid-19: systematic review and critical appraisal. BMJ. 2020 Apr 7; 369: m1328. https://doi.org/10.1136/bmj.m1328 PMID: 32265220
- Heaven WD. Hundreds of AI tools have been built to catch covid. None of them helped. MIT Technology Review [Internet]. 2021 Jul 30 [cited 2023 Jun 16]; Available from: https://www.technologyreview. com/2021/07/30/1030329/machine-learning-ai-failed-covid-hospital-diagnosis-pandemic/
- Hauser RG, Esserman D, Beste LA, Ong SY, Colomb DG, Bhargava A, et al. A Machine Learning Model to Successfully Predict Future Diagnosis of Chronic Myelogenous Leukemia With Retrospective Electronic Health Records Data. Am J Clin Pathol. 2021 Nov 8; 156(6):1142–8. <u>https://doi.org/10. 1093/ajcp/agab086 PMID: 34184028</u>
- Sendak M, Elish MC, Gao M, Futoma J, Ratliff W, Nichols M, et al. "The human body is a black box": supporting clinical decision-making with deep learning. In: Proceedings of the 2020 Conference on Fairness, Accountability, and Transparency. New York, NY, USA: Association for Computing Machinery; 2020. p. 99–109. (FAT\* '20).
- Abernethy A, Adams L, Barrett M, Bechtel C, Brennan P, Butte A, et al. The Promise of Digital Health: Then, Now, and the Future. NAM Perspect [Internet]. 2022 Jun 27;2022. Available from: <u>https://doi.org/10.31478/202206e PMID: 36177208</u>
- 20. Lavin A, Gilligan-Lee CM, Visnjic A, Ganju S, Newman D, Ganguly S, et al. Technology readiness levels for machine learning systems. Nat Commun. 2022 Oct 20; 13(1):6039. <u>https://doi.org/10.1038/s41467-022-33128-9 PMID: 36266298</u>

- D'Amour A, Heller K, Moldovan D, Adlam B, Alipanahi B, Beutel A, et al. Underspecification presents challenges for credibility in modern machine learning. J Mach Learn Res. 2022 Jan 1; 23(1):10237– 97.
- Benjamin R. Assessing risk, automating racism. Science. 2019 Oct 25; 366(6464):421–2. https://doi. org/10.1126/science.aaz3873 PMID: 31649182
- Cogburn CD. Culture, Race, and Health: Implications for Racial Inequities and Population Health. Milbank Q. 2019 Sep; 97(3):736–61. https://doi.org/10.1111/1468-0009.12411 PMID: 31512293
- 24. Assari S, Caldwell CH. Family Income at Birth and Risk of Attention Deficit Hyperactivity Disorder at Age 15: Racial Differences. Children [Internet]. 2019 Jan 14; 6(1). Available from: <a href="https://doi.org/10.3390/children6010010">https://doi.org/10.3390/children6010010</a> PMID: 30646634
- Brunner PM, Guttman-Yassky E. Racial differences in atopic dermatitis. Ann Allergy Asthma Immunol. 2019 May; 122(5):449–55. https://doi.org/10.1016/j.anai.2018.11.015 PMID: 30465859
- Murphy CC, Wallace K, Sandler RS, Baron JA. Racial Disparities in Incidence of Young-Onset Colorectal Cancer and Patient Survival. Gastroenterology. 2019 Mar; 156(4):958–65. <u>https://doi.org/10.1053/j.gastro.2018.11.060 PMID: 30521807</u>
- Obermeyer Z, Powers B, Vogeli C, Mullainathan S. Dissecting racial bias in an algorithm used to manage the health of populations. Science. 2019 Oct 25; 366(6464):447–53. <u>https://doi.org/10.1126/science.aax2342 PMID: 31649194</u>
- Okunlola OE, Lipnick MS, Batchelder PB, Bernstein M, Feiner JR, Bickler PE. Pulse Oximeter Performance, Racial Inequity, and the Work Ahead. Respir Care. 2022 Feb; 67(2):252–7. <u>https://doi.org/10.4187/respcare.09795 PMID: 34772785</u>
- Colvonen PJ, DeYoung PN, Bosompra NOA, Owens RL. Limiting racial disparities and bias for wearable devices in health science research. Sleep [Internet]. 2020 Oct 13; 43(10). Available from: <a href="https://doi.org/10.1093/sleep/zsaa159">https://doi.org/10.1093/sleep/zsaa159</a> PMID: 32893865
- Lane H, Sarkies M, Martin J, Haines T. Equity in healthcare resource allocation decision making: A systematic review. Soc Sci Med. 2017 Feb; 175:11–27. <u>https://doi.org/10.1016/j.socscimed.2016.12</u>. 012 PMID: 28043036
- Chen IY, Pierson E, Rose S, Joshi S, Ferryman K, Ghassemi M. Ethical Machine Learning in Healthcare. Annu Rev Biomed Data Sci. 2021 Jul; 4:123–44. https://doi.org/10.1146/annurev-biodatasci-092820-114757 PMID: 34396058
- Yu KH, Beam AL, Kohane IS. Artificial intelligence in healthcare. Nat Biomed Eng. 2018 Oct; 2 (10):719–31. https://doi.org/10.1038/s41551-018-0305-z PMID: 31015651
- Beam AL, Drazen JM, Kohane IS, Leong TY, Manrai AK, Rubin EJ. Artificial Intelligence in Medicine. N Engl J Med. 2023 Mar 30; 388(13):1220–1. <u>https://doi.org/10.1056/NEJMe2206291</u> PMID: 36988598
- 34. Finlayson SG, Subbaswamy A, Singh K, Bowers J, Kupke A, Zittrain J, et al. The Clinician and Dataset Shift in Artificial Intelligence. N Engl J Med. 2021 Jul 15; 385(3):283–6. <u>https://doi.org/10.1056/</u> NEJMc2104626 PMID: 34260843
- Nestor B, Hunter J, Kainkaryam R, Drysdale E, Inglis JB, Shapiro A, et al. Machine learning COVID-19 detection from wearables. Lancet Digit Health. 2023 Apr; 5(4):e182–4. <u>https://doi.org/10.1016/S2589-7500(23)00045-6</u> PMID: 36963907
- Ghassemi M, Mohamed S. Machine learning and health need better values. NPJ Digit Med. 2022 Apr 22; 5(1):51. https://doi.org/10.1038/s41746-022-00595-9 PMID: 35459793
- National Academy of Medicine. Artificial Intelligence in Health Care: The Hope, the Hype, the Promise, the Peril. 2023.
- Gichoya JW, Banerjee I, Bhimireddy AR, Burns JL, Celi LA, Chen LC, et al. AI recognition of patient race in medical imaging: a modelling study. Lancet Digit Health. 2022 Jun; 4(6):e406–14. <u>https://doi.org/10.1016/S2589-7500(22)00063-2 PMID: 35568690</u>
- 39. Yang MY, Kwak GH, Pollard T, Celi LA, Ghassemi M. Evaluating the Impact of Social Determinants on Health Prediction in the Intensive Care Unit. In: Proceedings of the 2023 AAAI/ACM Conference on AI, Ethics, and Society. New York, NY, USA: Association for Computing Machinery; 2023. p. 333–50. (AIES '23).
- Brönnimann S, Wintzer J. Climate data empathy. Wiley Interdisciplinary Reviews: Climate Change. 2019 Mar; 10(2):e559.
- **41.** Faghmous JH, Kumar V. A big data guide to understanding climate change: The case for theoryguided data science. Big Data. 2014 Sep; 2(3):155–63. https://doi.org/10.1089/big.2014.0026 PMID: 25276499

- 42. Institute of Medicine, Board on Health Sciences Policy, Committee on Understanding and Eliminating Racial and Ethnic Disparities in Health Care. Unequal Treatment: Confronting Racial and Ethnic Disparities in Health Care (with CD). National Academies Press; 2009. 432 p.
- Alberto IRI, Alberto NRI, Altinel Y, Blacker S, Binotti WW, Celi LA, et al. Who does the fairness in health AI community represent? [Internet]. medRxiv. 2023 [cited 2023 Jun 16].
  p. 2023.03.20.23287471. Available from: https://www.medrxiv.org/content/10.1101/2023.03.20. 23287471v1
- Mogensen L, Hu W. "A doctor who really knows ...": a survey of community perspectives on medical students and practitioners with disability. BMC Med Educ. 2019 Jul 29; 19(1):288.
- 45. Basu S, Berkowitz SA, Phillips RL, Bitton A, Landon BE, Phillips RS. Association of Primary Care Physician Supply With Population Mortality in the United States, 2005–2015. JAMA Intern Med. 2019 Apr 1; 179(4):506–14. https://doi.org/10.1001/jamainternmed.2018.7624 PMID: 30776056
- 46. Patrick-Lake B. Patient engagement in clinical trials: The Clinical Trials Transformation Initiative's leadership from theory to practical implementation. Clin Trials. 2018 Feb; 15(1\_suppl):19–22. https://doi.org/10.1177/1740774518755055 PMID: 29452519
- Domecq JP, Prutsky G, Elraiyah T, Wang Z, Nabhan M, Shippee N, et al. Patient engagement in research: a systematic review. BMC Health Serv Res. 2014 Feb 26; 14:89. https://doi.org/10.1186/ 1472-6963-14-89 PMID: 24568690
- Liberati A. Need to realign patient-oriented and commercial and academic research. Lancet. 2011 Nov 19; 378(9805):1777–8.
- Sacristán JA, Aguarón A, Avendaño-Solá C, Garrido P, Carrión J, Gutiérrez A, et al. Patient involvement in clinical research: why, when, and how. Patient Prefer Adherence. 2016 Apr 27; 10:631–40. https://doi.org/10.2147/PPA.S104259 PMID: 27175063
- Silvola S, Restelli U, Bonfanti M, Croce D. Co-Design as Enabling Factor for Patient-Centred Healthcare: A Bibliometric Literature Review. Clinicoecon Outcomes Res. 2023 May 17; 15:333–47. https:// doi.org/10.2147/CEOR.S403243 PMID: 37220481
- Malekinejad M, Horvath H, Snyder H, Brindis CD. The discordance between evidence and health policy in the United States: the science of translational research and the critical role of diverse stakeholders. Health Res Policy Syst. 2018 Aug 16; 16(1):81. <u>https://doi.org/10.1186/s12961-018-0336-7</u> PMID: 30115085
- 52. Aboab J, Celi LA, Charlton P, Feng M, Ghassemi M, Marshall DC, et al. A "datathon" model to support cross-disciplinary collaboration. Sci Transl Med. 2016 Apr 6; 8(333):333ps8.
- 53. NeurIPS 2021 [Internet]. [cited 2023 Jul 19]. Available from: <u>https://nips.cc/Conferences/2021/</u> DatasetsBenchmarks/AcceptedPapers
- Ng MY, Youssef A, Miner AS, Sarellano D, Long J, Larson DB, et al. Perceptions of Data Set Experts on Important Characteristics of Health Data Sets Ready for Machine Learning: A Qualitative Study. JAMA Netw Open. 2023 Dec 1; 6(12):e2345892. <u>https://doi.org/10.1001/jamanetworkopen.2023</u>. 45892 PMID: 38039004
- 55. Nyström ME, Karltun J, Keller C, Andersson Gäre B. Collaborative and partnership research for improvement of health and social services: researcher's experiences from 20 projects. Health Res Policy Syst. 2018 May 30; 16(1):46. https://doi.org/10.1186/s12961-018-0322-0 PMID: 29843735
- 56. Kaushal A, Altman R, Langlotz C. Geographic Distribution of US Cohorts Used to Train Deep Learning Algorithms. JAMA. 2020 Sep 22; 324(12):1212–3. <u>https://doi.org/10.1001/jama.2020.12067</u> PMID: 32960230
- McDermott MBA, Wang S, Marinsek N, Ranganath R, Foschini L, Ghassemi M. Reproducibility in machine learning for health research: Still a ways to go. Sci Transl Med [Internet]. 2021 Mar 24; 13 (586). Available from: http://dx.doi.org/10.1126/scitranslmed.abb1655
- Paterson R, MacLeod DC, Thetford D, Beattie A, Graham C, Lam S, et al. Prediction of in-hospital mortality and length of stay using an early warning scoring system: clinical audit. Clin Med. 2006 May-Jun; 6(3):281–4. https://doi.org/10.7861/clinmedicine.6-3-281 PMID: 16826863
- 59. Stone K, Zwiggelaar R, Jones P, Mac Parthaláin N. A systematic review of the prediction of hospital length of stay: Towards a unified framework. PLOS Digit Health. 2022 Apr; 1(4):e0000017. https://doi. org/10.1371/journal.pdig.0000017 PMID: 36812502
- 60. Aoyama K, D'Souza R, Pinto R, Ray JG, Hill A, Scales DC, et al. Risk prediction models for maternal mortality: A systematic review and meta-analysis. PLoS One. 2018 Dec 4; 13(12):e0208563. <u>https://doi.org/10.1371/journal.pone.0208563 PMID: 30513118</u>
- Ruiz JI, Nuhu K, McDaniel JT, Popoff F, Izcovich A, Criniti JM. Inequality as a Powerful Predictor of Infant and Maternal Mortality around the World. PLoS One. 2015 Oct 21; 10(10):e0140796. <u>https://doi.org/10.1371/journal.pone.0140796 PMID: 26488170</u>

- Wasserman J, Palmer RC, Gomez MM, Berzon R, Ibrahim SA, Ayanian JZ. Advancing Health Services Research to Eliminate Health Care Disparities. Am J Public Health. 2019 Jan; 109(S1):S64–9. https://doi.org/10.2105/AJPH.2018.304922 PMID: 30699021
- 63. Wang F. Why Public Health Needs GIS: A Methodological Overview. Ann GIS. 2020; 26(1):1–12. https://doi.org/10.1080/19475683.2019.1702099 PMID: 32547679
- 64. Aim-ahead [Internet]. [cited 2023 Dec 25]. Available from: https://datascience.nih.gov/artificialintelligence/aim-ahead
- 65. Bridge to Artificial Intelligence (Bridge2AI) [Internet]. [cited 2023 Dec 25]. Available from: https:// commonfund.nih.gov/bridge2ai
- Harnessing Data Science for Health Discovery and Innovation in Africa (DS-I Africa) [Internet]. [cited 2023 Dec 25]. Available from: https://commonfund.nih.gov/AfricaData
- 67. Gates foundation celebrates 20 years of "grand challenges" with new investments and a call to make R&D breakthroughs available more quickly and equitably [Internet]. Bill & Melinda Gates Foundation. [cited 2023 Dec 25]. Available from: https://www.gatesfoundation.org/ideas/media-center/press-releases/2023/10/grand-challenges-ai-equity-womens-health
- **68.** Bolukbasi E. Listening to scientists from across Africa to develop our next Mental Health Data Prize [Internet]. Wellcome Trust. 2023 [cited 2023 Dec 25]. Available from: <u>https://wellcome.org/news/</u>listening-scientists-across-africa-develop-our-next-mental-health-data-prize
- 69. Manda MI, Backhouse J. Addressing trust, security and privacy concerns in e-government integration, interoperability and information sharing through policy: a case of South Africa. 2016 [cited 2023 Dec 25]; Available from: https://aisel.aisnet.org/confirm2016/67/
- 70. Dhagarra D, Goswami M, Kumar G. Impact of Trust and Privacy Concerns on Technology Acceptance in Healthcare: An Indian Perspective. Int J Med Inform. 2020 Sep; 141:104164. <u>https://doi.org/10. 1016/j.ijmedinf.2020.104164 PMID: 32593847</u>
- 71. Restrepo D, Quion J, Vásquez-Venegas C, Villanueva C, Anthony Celi L, Nakayama LF. A scoping review of the landscape of health-related open datasets in Latin America. PLOS Digit Health. 2023 Oct; 2(10):e0000368. https://doi.org/10.1371/journal.pdig.0000368 PMID: 37878549
- 72. Rostamzadeh N, Mincu D, Roy S, Smart A, Wilcox L, Pushkarna M, et al. Healthsheet: Development of a Transparency Artifact for Health Datasets. In: Proceedings of the 2022 ACM Conference on Fairness, Accountability, and Transparency. New York, NY, USA: Association for Computing Machinery; 2022. p. 1943–61. (FAccT '22).
- 73. Gebru T, Morgenstern J, Vecchione B, Vaughan JW, Wallach H, Iii HD, et al. Datasheets for datasets. Commun ACM. 2021 Nov 19; 64(12):86–92.
- 74. Simon GE, Shortreed SM, Coley RY, Penfold RB, Rossom RC, Waitzfelder BE, et al. Assessing and Minimizing Re-identification Risk in Research Data Derived from Health Care Records. EGEMS (Wash DC). 2019 Mar 29; 7(1):6. https://doi.org/10.5334/egems.270 PMID: 30972355
- Xiao, Y., Lim, S., Pollard, T. J. & Ghassemi, M. In the Name of Fairness: Assessing the Bias in Clinical Record De-identification. Proceedings of the 2023 Conference on Fairness, Accountability, and Transparency (2023).
- 76. Johnson AEW, Pollard TJ, Shen L, Lehman LWH, Feng M, Ghassemi M, et al. MIMIC-III, a freely accessible critical care database. Sci Data. 2016 May 24; 3:160035. <u>https://doi.org/10.1038/sdata.2016.35 PMID: 27219127</u>
- 77. de Kok JWTM, de la Hoz MÁA, de Jong Y, Brokke V, Elbers PWG, Thoral P, et al. A guide to sharing open healthcare data under the General Data Protection Regulation. Sci Data. 2023 Jun 24; 10 (1):404. https://doi.org/10.1038/s41597-023-02256-2 PMID: 37355751
- Moody GB, Mark RG. A database to support development and evaluation of intelligent intensive care monitoring. In: Computers in Cardiology 1996. ieeexplore.ieee.org; 1996. p. 657–60.
- 79. Wilkinson MD, Dumontier M, Aalbersberg IJJ, Appleton G, Axton M, Baak A, et al. The FAIR Guiding Principles for scientific data management and stewardship. Sci Data. 2016 Mar 15; 3:160018. <u>https:// doi.org/10.1038/sdata.2016.18 PMID: 26978244</u>
- Callender T, van der Schaar M. Automated machine learning as a partner in predictive modelling. Lancet Digit Health. 2023 May; 5(5):e254–6. https://doi.org/10.1016/S2589-7500(23)00054-7 PMID: 37100541
- Hripcsak G, Duke JD, Shah NH, Reich CG, Huser V, Schuemie MJ, et al. Observational Health Data Sciences and Informatics (OHDSI): Opportunities for Observational Researchers. Stud Health Technol Inform. 2015; 216:574–8. https://doi.org/10.1038/psp.2013.52 PMID: 26262116
- Mitchell M, Wu S, Zaldivar A, Barnes P, Vasserman L, Hutchinson B, et al. Model Cards for Model Reporting. In: Proceedings of the Conference on Fairness, Accountability, and Transparency. New York, NY, USA: Association for Computing Machinery; 2019. p. 220–9. (FAT\* '19).

- Wiens J, Saria S, Sendak M, Ghassemi M, Liu VX, Doshi-Velez F, et al. Do no harm: a roadmap for responsible machine learning for health care. Nat Med. 2019 Sep; 25(9):1337–40. <u>https://doi.org/10. 1038/s41591-019-0548-6 PMID: 31427808</u>
- Hayat, N., Geras, K. J. & Shamout, F. E. MedFuse: Multi-modal fusion with clinical time-series data and chest X-ray images. In Machine Learning for Healthcare Conference (pp. 479–503). PMLR.
- Zhang H, Dullerud N, Roth K, Oakden-Rayner L, Pfohl S, Ghassemi M. Improving the Fairness of Chest X-ray Classifiers. In: Flores G, Chen GH, Pollard T, Ho JC, Naumann T, editors. Proceedings of the Conference on Health, Inference, and Learning. PMLR; 07–08 Apr 2022. p. 204–33. (Proceedings of Machine Learning Research; vol. 174).
- Iancu A, Leb I, Prokosch HU, Rödle W. Machine learning in medication prescription: A systematic review. Int J Med Inform. 2023 Dec; 180:105241. https://doi.org/10.1016/j.ijmedinf.2023.105241 PMID: 37939541
- Rodziewicz TL, Houseman B, Hipskind JE. Medical Error Reduction and Prevention. In: StatPearls. Treasure Island (FL): StatPearls Publishing; 2023.
- 88. FitzGerald C, Hurst S. Implicit bias in healthcare professionals: a systematic review. BMC Med Ethics. 2017 Mar 1; 18(1):19. https://doi.org/10.1186/s12910-017-0179-8 PMID: 28249596
- Schoenthaler A, Williams N. Looking Beneath the Surface: Racial Bias in the Treatment and Management of Pain. JAMA Netw Open. 2022 Jun 1; 5(6):e2216281. <u>https://doi.org/10.1001/jamanetworkopen.2022.16281 PMID: 35679049</u>
- 90. Pierson E, Cutler DM, Leskovec J, Mullainathan S, Obermeyer Z. An algorithmic approach to reducing unexplained pain disparities in underserved populations. Nat Med. 2021 Jan; 27(1):136–40. <u>https://</u> doi.org/10.1038/s41591-020-01192-7 PMID: 33442014
- Tan KR, Seng JJB, Kwan YH, Chen YJ, Zainudin SB, Loh DHF, et al. Evaluation of Machine Learning Methods Developed for Prediction of Diabetes Complications: A Systematic Review. J Diabetes Sci Technol. 2023 Mar; 17(2):474–89. https://doi.org/10.1177/19322968211056917 PMID: 34727783
- Garbin C, Marques N, Marques O. Machine learning for predicting opioid use disorder from healthcare data: A systematic review. Comput Methods Programs Biomed. 2023 Jun; 236:107573. <u>https://doi.org/10.1016/j.cmpb.2023.107573 PMID</u>: 37148670
- Islam KR, Prithula J, Kumar J, Tan TL, Reaz MBI, Sumon MSI, et al. Machine Learning-Based Early Prediction of Sepsis Using Electronic Health Records: A Systematic Review. J Clin Med Res [Internet]. 2023 Aug 30; 12(17). Available from: https://doi.org/10.3390/jcm12175658 PMID: 37685724
- 94. Wawira Gichoya J, McCoy LG, Celi LA, Ghassemi M. Equity in essence: a call for operationalising fairness in machine learning for healthcare. BMJ Health Care Inform [Internet]. 2021 Apr; 28(1). Available from: http://dx.doi.org/10.1136/bmjhci-2020-100289
- 95. Adam H, Yang MY, Cato K, Baldini I, Senteio C, Celi LA, et al. Write it like you see it: Detectable differences in clinical notes by race lead to differential model recommendations. In: Proceedings of the 2022 AAAI/ACM Conference on AI, Ethics, and Society [Internet]. New York, NY, USA: ACM; 2022. Available from: https://dl.acm.org/doi/10.1145/3514094.3534203
- 96. Bhanot K, Soares IB, Wei D, Zeng J, Bennett K. Stress-testing Bias Mitigation Algorithms to Understand Fairness Vulnerabilities. In: AAAI/ACM Conference on AI, Ethics, and Society [Internet]. 2021 [cited 2023 Jul 20]. Available from: https://research.ibm.com/publications/stress-testing-biasmitigation-algorithms-to-understand-fairness-vulnerabilities
- Schrouff J, Harris N, Koyejo O, Alabdulmohsin I, Schnider E, Opsahl-Ong K, et al. Diagnosing failures of fairness transfer across distribution shift in real-world medical settings. Advances in Neural Information Processing Systems, 35, 19304–19318.
- Parbhoo S, Wawira Gichoya J, Celi LA, de la Hoz MÁA, for MIT Critical Data. Operationalising fairness in medical algorithms. BMJ Health Care Inform [Internet]. 2022 Jun; 29(1). Available from: <u>http://dx.</u> doi.org/10.1136/bmjhci-2022-100617
- 99. Beaulieu-Jones B, Finlayson SG, Chivers C, Chen I, McDermott M, Kandola J, et al. Trends and Focus of Machine Learning Applications for Health Research. JAMA Netw Open. 2019 Oct 2; 2(10): e1914051. https://doi.org/10.1001/jamanetworkopen.2019.14051 PMID: 31651969
- 100. Zhang A, Xing L, Zou J, Wu JC. Shifting machine learning for healthcare from development to deployment and from models to data. Nat Biomed Eng. 2022 Dec; 6(12):1330–45. <u>https://doi.org/10.1038/</u> s41551-022-00898-y PMID: 35788685
- 101. Nestor B, McDermott MBA, Boag W, Berner G, Naumann T, Hughes MC, et al. Feature Robustness in Non-stationary Health Records: Caveats to Deployable Model Performance in Common Clinical Machine Learning Tasks. In: Doshi-Velez F, Fackler J, Jung K, Kale D, Ranganath R, Wallace B, et al., editors. Proceedings of the 4th Machine Learning for Healthcare Conference. PMLR; 09–10 Aug 2019. p. 381–405. (Proceedings of Machine Learning Research; vol. 106).

- 102. Kulkarni V, Gawali M, Kharat A. Key Technology Considerations in Developing and Deploying Machine Learning Models in Clinical Radiology Practice. JMIR Med Inform. 2021 Sep 9; 9(9):e28776. https://doi.org/10.2196/28776 PMID: 34499049
- 103. Ahmad T, Desai NR, Yamamoto Y, Biswas A, Ghazi L, Martin M, et al. Alerting Clinicians to 1-Year Mortality Risk in Patients Hospitalized With Heart Failure: The REVEAL-HF Randomized Clinical Trial. JAMA Cardiol. 2022 Sep 1; 7(9):905–12. <u>https://doi.org/10.1001/jamacardio.2022.2496</u> PMID: 35947362
- 104. Verma AA, Pou-Prom C, McCoy LG, Murray J, Nestor B, Bell S, et al. Developing and Validating a Prediction Model For Death or Critical Illness in Hospitalized Adults, an Opportunity for Human-Computer Collaboration. Crit Care Explor. 2023 May; 5(5):e0897. https://doi.org/10.1097/CCE. 00000000000897 PMID: 37151895
- 105. Guo C, Ashrafian H, Ghafur S, Fontana G, Gardner C, Prime M. Challenges for the evaluation of digital health solutions—A call for innovative evidence generation approaches. npj Digital Medicine. 2020 Aug 27; 3(1):1–14. https://doi.org/10.1038/s41746-020-00314-2 PMID: 32904379
- 106. Sendak MP, Ratliff W, Sarro D, Alderton E, Futoma J, Gao M, et al. Real-World Integration of a Sepsis Deep Learning Technology Into Routine Clinical Care: Implementation Study. JMIR Med Inform. 2020 Jul 15; 8(7):e15182. https://doi.org/10.2196/15182 PMID: 32673244
- 107. Sambasivan N, Kapania S, Highfill H, Akrong D, Paritosh P, Aroyo LM. "Everyone wants to do the model work, not the data work": Data Cascades in High-Stakes Al. In: Proceedings of the 2021 CHI Conference on Human Factors in Computing Systems. New York, NY, USA: Association for Computing Machinery; 2021. p. 1–15. (CHI '21).
- Paleyes A, Urma RG, Lawrence ND. Challenges in Deploying Machine Learning: A Survey of Case Studies. ACM Comput Surv. 2022 Dec 7; 55(6):1–29.
- 109. Glasgow RE, Vogt TM, Boles SM. Evaluating the public health impact of health promotion interventions: the RE-AIM framework. Am J Public Health. 1999 Sep; 89(9):1322–7. <u>https://doi.org/10.2105/ajph.89.9.1322</u> PMID: 10474547
- Song Z, Kannan S, Gambrel RJ, Marino M, Vaduganathan M, Clapp MA, et al. Physician Practice Pattern Variations in Common Clinical Scenarios Within 5 US Metropolitan Areas. JAMA Health Forum. 2022 Jan; 3(1):e214698. https://doi.org/10.1001/jamahealthforum.2021.4698 PMID: 35977237
- 111. Albanowski K, Burdick KJ, Bonafide CP, Kleinpell R, Schlesinger JJ. Ten Years Later, Alarm Fatigue Is Still a Safety Concern. AACN Adv Crit Care. 2023 Sep 15; 34(3):189–97. <u>https://doi.org/10.4037/aacnacc2023662 PMID: 37644627</u>
- 112. Kostick-Quenet KM, Gerke S. Al in the hands of imperfect users. NPJ Digit Med. 2022 Dec 28; 5 (1):197. https://doi.org/10.1038/s41746-022-00737-z PMID: 36577851
- 113. Gaube S, Suresh H, Raue M, Merritt A, Berkowitz SJ, Lermer E, et al. Do as Al say: susceptibility in deployment of clinical decision-aids. NPJ Digit Med. 2021 Feb 19; 4(1):31. <u>https://doi.org/10.1038/</u> s41746-021-00385-9 PMID: 33608629
- 114. Goddard K, Roudsari A, Wyatt JC. Automation bias: a systematic review of frequency, effect mediators, and mitigators. J Am Med Inform Assoc. 2012 Jan-Feb; 19(1):121–7. https://doi.org/10.1136/ amiajnl-2011-000089 PMID: 21685142
- 115. Grote T, Berens P. On the ethics of algorithmic decision-making in healthcare. J Med Ethics. 2020 Mar; 46(3):205–11. https://doi.org/10.1136/medethics-2019-105586 PMID: 31748206
- 116. Dietvorst BJ, Simmons JP, Massey C. Algorithm aversion: people erroneously avoid algorithms after seeing them err. J Exp Psychol Gen. 2015 Feb; 144(1):114–26. <u>https://doi.org/10.1037/xge0000033</u> PMID: 25401381
- 117. Chromik J, Klopfenstein SAI, Pfitzner B, Sinno ZC, Arnrich B, Balzer F, et al. Computational approaches to alleviate alarm fatigue in intensive care medicine: A systematic literature review. Front Digit Health. 2022 Aug 16; 4:843747. https://doi.org/10.3389/fdgth.2022.843747 PMID: 36052315
- 118. Adam H, Balagopalan A, Alsentzer E, Christia F, Ghassemi M. Mitigating the impact of biased artificial intelligence in emergency decision-making. Commun Med. 2022 Nov 21; 2(1):149. <u>https://doi.org/10. 1038/s43856-022-00214-4 PMID: 36414774</u>
- 119. Lu J. Will medical technology deskill doctors? International Education Studies [Internet]. 2016; Available from: https://hub.hku.hk/handle/10722/231367
- 120. Madras D, Pitassi T, Zemel R. Predict responsibly: improving fairness and accuracy by learning to defer. Adv Neural Inf Process Syst [Internet]. 2018; Available from: https://proceedings.neurips.cc/ paper/2018/hash/09d37c08f7b129e96277388757530c72-Abstract.html
- 121. Mozannar H, Satyanarayan A, Sontag D. Teaching Humans When to Defer to a Classifier via Exemplars. AAAI. 2022 Jun 28; 36(5):5323–31.

- 122. Singhal K, Tu T, Gottweis J, Sayres R, Wulczyn E, Hou L, et al. Towards Expert-Level Medical Question Answering with Large Language Models [Internet]. arXiv [cs.CL]. 2023. Available from: <u>http://arxiv.org/abs/2305.09617</u>
- 123. Otokiti AU, Ozoude MM, Williams KS, Sadiq-Onilenla RA, Ojo SA, Wasarme LB, et al. The Need to Prioritize Model-Updating Processes in Clinical Artificial Intelligence (AI) Models: Protocol for a Scoping Review. JMIR Res Protoc. 2023 Feb 16; 12:e37685. <u>https://doi.org/10.2196/37685</u> PMID: 36795464
- 124. Wong A, Otles E, Donnelly JP, Krumm A, McCullough J, DeTroyer-Cooley O, et al. External Validation of a Widely Implemented Proprietary Sepsis Prediction Model in Hospitalized Patients. JAMA Intern Med. 2021 Aug 1; 181(8):1065–70. https://doi.org/10.1001/jamainternmed.2021.2626 PMID: 34152373
- 125. Greenhalgh T, Humphrey C, Hughes J, Macfarlane F, Butler C, Pawson R. How do you modernize a health service? A realist evaluation of whole-scale transformation in london. Milbank Q. 2009 Jun; 87 (2):391–416. https://doi.org/10.1111/j.1468-0009.2009.00562.x PMID: 19523123
- 126. Jack K. What is realist evaluation? Evid Based Nurs [Internet]. 2022 Aug 19; Available from: https:// doi.org/10.1136/ebnurs-2022-103608 PMID: 35985802
- 127. Beam K, Sharma P, Levy P, Beam AL. Artificial intelligence in the neonatal intensive care unit: the time is now. J Perinatol [Internet]. 2023 Jul 13; Available from: https://doi.org/10.1038/s41372-023-01719-z PMID: 37443271
- 128. Mann KD, Good NM, Fatehi F, Khanna S, Campbell V, Conway R, et al. Predicting Patient Deterioration: A Review of Tools in the Digital Hospital Setting. J Med Internet Res. 2021 Sep 30; 23(9):e28209. https://doi.org/10.2196/28209 PMID: 34591017
- 129. Wang D, Mo J, Zhou G, Xu L, Liu Y. An efficient mixture of deep and machine learning models for COVID-19 diagnosis in chest X-ray images. PLoS One. 2020 Nov 17; 15(11):e0242535. <u>https://doi.org/10.1371/journal.pone.0242535 PMID: 33201919</u>
- 130. Krusche M, Callhoff J, Knitza J, Ruffer N. Diagnostic accuracy of a large language model in rheumatology: comparison of physician and ChatGPT-4. Rheumatol Int [Internet]. 2023 Sep 24; Available from: https://doi.org/10.1007/s00296-023-05464-6 PMID: 37742280
- 131. Burger R, Christian C. Access to health care in post-apartheid South Africa: availability, affordability, acceptability. Health Econ Policy Law. 2020 Jan; 15(1):43–55. <u>https://doi.org/10.1017/S1744133118000300 PMID: 29996951</u>
- 132. Sekimitsu S, Zebardast N. Glaucoma and machine learning: A call for increased diversity in data. Ophthalmol Glaucoma. 2021; 4(4):339–42. <u>https://doi.org/10.1016/j.ogla.2021.03.002</u> PMID: 33879422