Title
Artificial intelligence in gastroenterology and hepatology: how to advance clinical practice while ensuring health equity.

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Artificial intelligence in gastroenterology and hepatology: how to advance clinical practice while ensuring health equity

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ABSTRACT
Artificial intelligence (AI) and machine learning (ML) systems are increasingly used in medicine to improve clinical decision-making and healthcare delivery. In gastroenterology and hepatology, studies have explored a myriad of opportunities for AI/ML applications which are already making the transition to bedside. Despite these advances, there is a risk that biases and health inequities can be introduced or exacerbated by these technologies. If unrecognised, these technologies could generate or worsen systematic racial, ethnic and sex disparities when deployed on a large scale. There are several mechanisms through which AI/ML could contribute to health inequities in gastroenterology and hepatology, including diagnosis of oesophageal cancer, management of inflammatory bowel disease (IBD), liver transplantation, colorectal cancer screening and many others. This review adapts a framework for ethical AI/ML development and application to gastroenterology and hepatology such that clinical practice is advanced while minimising bias and optimising health equity.

Key messages
- Artificial intelligence (AI) and machine learning (ML) systems are increasingly used in medicine to improve clinical decision-making and healthcare delivery.
- In gastroenterology and hepatology, studies have explored a myriad of opportunities for AI/ML applications which are already making the transition to bedside.
- Despite these advances, there is a risk that biases and health inequities can be introduced or exacerbated by these technologies. If unrecognised, these technologies could generate or worsen systematic racial, ethnic and sex disparities when deployed on a large scale.
- There are several mechanisms through which AI/ML could contribute to health inequities in gastroenterology and hepatology, including diagnosis of oesophageal cancer, management of inflammatory bowel disease (IBD), liver transplantation, colorectal cancer screening and many others.
- This review adapts a framework for ethical AI/ML development and application to gastroenterology and hepatology such that clinical practice is advanced while minimising bias and optimising health equity.

INTRODUCTION: ARTIFICIAL INTELLIGENCE AND HEALTH EQUITY
Artificial intelligence (AI) and machine learning (ML) technologies can leverage massive amounts of data for predictive modelling in a wide variety of fields and are increasingly used to inform complex decision-making and clinical processes in healthcare.1 Examples include computer vision-assisted mammograms to improve breast cancer detection,2 models that predict respiratory decompensation in patients with COVID-193 and AI tools which predict length of stay, facilitate resource allocation and lower healthcare costs.4
In gastroenterology and hepatology, opportunities for AI/ML implementation are burgeoning. Recent studies have explored AI applications such as computer-aided detection (CADe) for diagnosis of premalignant and malignant GI lesions, prediction of treatment response in patients with inflammatory bowel disease (IBD), histopathological analysis of biopsy specimens, assessment of liver fibrosis severity in chronic liver disease, models for liver transplant allocation and others.5-8 AI-based systems are increasingly making the transition from research to bedside and have the potential to revolutionise patient care. However, these advances must be matched by corresponding regulatory and ethical frameworks developed by the Food and Drug Administration (FDA) and other agencies that oversee the intended and unintended consequences of their use.9 Concerns have already been raised regarding the biases and health inequities that can be introduced or amplified when applying computer algorithms in healthcare.10 For instance, a commercial algorithm applied to approximately 200 million patients in the USA was racially biased—white patients were preferentially enrolled in ‘high-risk care management programmes’ compared with black patients with similar risk scores, resulting in fewer healthcare dollars spent on black patients.11 Another study demonstrated that an ML algorithm that predicts intensive care unit mortality and 30-day psychiatric readmission rates had poorer predictive performance for women and patients with public insurance.12
Prior to deploying and scaling AI/ML tools, it is critical to ensure that the risk of bias is minimised...
and opportunities to promote health equity are amplified. In this work, we identify areas in gastroenterology and hepatology where algorithms could exacerbate disparities, and offer potential areas of opportunity for advancing health equity through AI/ML. For the purposes of this paper, health equity is centred on distributive justice to eliminate systematic racial, ethnic and sex disparities.13

**FIVE KEY MECHANISMS THROUGH WHICH AI/ML CAN CONTRIBUTE TO HEALTH INEQUITIES**

Early efforts to promote responsible and ethical applications of AI and ML in clinical medicine have revealed several mechanisms through which algorithms can introduce bias and exacerbate health inequities (figure 1).10 14 It is essential for researchers and clinicians in gastroenterology and hepatology to understand these mechanisms as new technologies are being applied to our field.

The first theme is disparities in clinical or research problem selection14—research questions often target concerns in majority populations due to unequal funding availability and/or interest in the problem by industry, researchers, funders and grant review committees. As a result, we see critical racial, ethnic and sex disparities in the research problems that are prioritised and funded in AI/ML.14 15

The second theme is bias in data collection,14 where collected data may capture a disproportionate share of one population group over another. This can result in algorithms that are not widely generalisable,14 16–19 especially for individuals from traditionally under-represented and marginalised groups that are not commonly or appreciably represented in research databases.

The third theme is bias due to variable selection.14 Variables and outcome measures may appear unbiased on initial evaluation even though they are proxies for, or confounded by, explicit or implicit biases against under-represented or marginalised groups.

The fourth theme is bias in algorithm development.14 In this case, the assumptions made and used by the research team lead to inherently biased models or models that are overfitted to narrow training data.13 15 17 Additionally, the performance metrics of AI/ML model training, such as area under the curve, are not inherently optimal for equitable performance in diverse populations.14 20

Finally, inequities may result from post-deployment considerations.14 Even a potentially unbiased AI tool may lead to biased behaviour when deployed in the clinical setting. It is important to consider (1) how a tool will perform in a disease that has different conditional distributions in a population and (2) the potentially negative human–computer interactions that may occur. For instance, providers may follow an AI/ML-generated treatment recommendation when it confirms their biased beliefs but disregard treatment recommendations that do not conform to their beliefs.14 21 The impacts of the algorithm should be evaluated in population subgroups to assess differences in clinical behaviour, performance and outcomes by sociodemographic factors, rather than at the population level alone.14

**AI IN GASTROENTEROLOGY AND HEPATOLOGY: IMPACTS ON HEALTH EQUITY**

We have chosen specific clinical examples from the literature to illustrate the specific ways existing AI/ML algorithms may already exacerbate bias and inequities within the fields of gastroenterology and hepatology (table 1).

**Oesophageal cancer**

Prevention and early recognition of oesophageal cancer is an area in which AI may hold particular promise, and there has been meaningful research progress in this area. In the USA, the vast majority of oesophageal cancer research focuses on technologies (with and without AI) to improve the early identification and treatment of Barrett’s oesophagus and oesophageal adenocarcinoma (OAC).22–26 Unfortunately, this emphasis on OAC in the USA primarily benefits white populations, who have the highest incidence and mortality from OAC.27

Oesophageal squamous cell carcinoma (OSCC) is more common than OAC and has a higher incidence and mortality in non-white populations in the USA and worldwide.28 29 Specifically, black individuals in the USA have the highest incidence of OSCC at 4.9 per 100 000 people, followed by Asians at 1.9 per 100 000, and white individuals with the lowest rates at 1.4 per 100 000. These rates are higher than overall OAC rates and OAC rates among non-white individuals in the USA: 2.3 per 100 000 for white individuals, compared with 0.5 per 100 000 in black and Asian individuals.28 Furthermore, AI research in OSCC has largely been performed in Asian countries, and thus, it is uncertain whether findings may be generalisable to black individuals or other population subgroups globally.30–34

This research disparity may be due to a clinical or research problem selection bias (theme 1); both researchers and funders should work to ensure more equity in problem selection. AI-based tools for early recognition of oesophageal cancer and precursor lesions have the potential to save many lives, but in the current state will largely impact white patients and not individuals from under-represented groups. It is imperative that we consider inclusive AI/ML research questions to avoid preferential development of technologies that consistently benefit one group over others.

Figure 1 Mechanisms through which AI contributes to health inequities. Adapted from Chen et al.14 AI, artificial intelligence.
Inflammatory bowel disease

In IBD, AI/ML computer vision tools have been developed for endoscopic assessment of disease severity, to distinguish colitis from neoplasia, and to differentiate sporadic adenomas from non-neoplastic lesions.34 35 AI algorithms have also been trained to predict treatment response and assess risk of disease recurrence.36–39 AI has the potential to play an important role in IBD treatment decisions by predicting response earlier in the treatment course and guiding personalised therapy choices.

However, many AI/ML models developed in IBD have been created in largely white populations. For example, one study used AI to predict future corticosteroid use and hospitalisation in patients with IBD from a cohort of 20,368 patients at the Veterans Health Administration (VA).36 The authors concluded that their model had the potential to predict IBD flares, improve patient outcomes and reduce healthcare costs. They also noted that their algorithm would be easy to implement at the point of care to individualise and tailor therapies for individual patients. The population that was used to derive the model was 93% male which may make predictions for female patients with IBD less relevant. The algorithm also included race as a predictor, though the dataset was racially skewed: the study population was 70% white, though women were more adequately represented (57.1%). The authors concluded that their model could be used to predict future corticosteroid use and hospitalisation in patients with IBD from a cohort of 20,368 patients at the Veterans Health Administration (VA).36

In the VA study, the proportion of the population that was Hispanic/Latinx or South and East Asian was not included, despite the fact that these groups comprise an increasingly large share of the populations with IBD. While this study may be beneficial to the patient population served by the VA, it suggests that even in very large cohorts, there may be entrenched patterns of bias in data collection (theme 2): algorithms that do not include the rich diversity of patients with IBD can result in biased systems, care and outcomes, particularly if extrapolated to the general population.

Liver transplantation

There are numerous opportunities for AI/ML applications in hepatology, including the assessment of hepatic fibrosis progression, detection of non-alcoholic fatty liver disease, identification of patients at risk of hepatocellular carcinoma (HCC) and optimisation of organ transplant protocols.40–43 As we explore the complex and opaque nature of emerging AI clinical prediction tools, it is important to recognise that bias can be encoded even in conventional prediction models, including simple, rule-based algorithms.

Prior to the adoption of the Model for End-Stage Liver Disease (MELD) score in 2002, the liver allocation process was fraught with variability, subjectivity and opportunities for manipulation, which resulted in inequities.44 To address these shortcomings and standardise the organ allocation process, the United Network for Organ Sharing turned to the MELD—an algorithmic model which predicts 3-month survival rates in patients with cirrhosis—as a way to more fairly prioritise patients for liver transplantation.45 Variables included in the model appear to be objective laboratory values—bilirubin, creatinine, international normalised ratio and sodium. Creatinine however underestimates renal dysfunction in women, leading to lower MELD scores compared with men with similar disease severity. This underestimation negatively impacts equitable organ allocation for liver transplant.

A similar example occurs with MELD exception points—a system where patients with certain conditions that confer excess risk beyond that captured by the laboratory variables that comprise MELD (such as HCC) may accumulate points and advance their position on the transplant waitlist.46 Review of data from Organ Procurement and Transplantation Network (OPTN) registries shows that at similar listing priority, patients with MELD exception points are less likely to die on the waitlist, more likely to receive a transplant and less likely to be women.47 48 49 Part of this discrepancy is because HCC—the indication for MELD exception points in approximately 70% of

### Table 1  Types of bias observed in artificial intelligence (AI) in clinical medicine

<table>
<thead>
<tr>
<th>Theme</th>
<th>Definition</th>
<th>Examples</th>
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| Problem selection | Differential research priority and funding for issues that affect marginalised groups. | - AI has been used extensively to detect Barrett’s oesophagus and oesophageal adenocarcinoma, which mainly affects white individuals.  
- In contrast, AI applications in oesophageal squamous carcinoma—which is more prevalent in underserved populations—are under-researched. |
| Data collection | Inadequate representation of underserved groups in training datasets results in biased algorithms that yield inaccurate outputs for these subgroups. | - A model trained on Veteran’s Health Administration electronic database to predict IBD flares may generate incorrect predictions for non-white populations who are under-represented in the training dataset. |
| Variable selection | Seemingly objective predictor and outcome variables that are included in a model may be confounded by or proxies for factors that lead to biased results. | - MELD exception points may appropriately prioritise patients with HCC on the transplant waitlist; however, this is confounded by the increased prevalence of HCC in men which leads to lower transplant rates for women.  
- The inclusion of serum creatinine in the MELD leads to lower scores and transplant priority for women as serum creatinine underestimates renal dysfunction in women. |
| Algorithm development | Models are developed to recreate patterns in the training dataset and may not account for systemic biases. | - Racial and ethnic minorities are less likely to be referred for liver transplant and more likely to be offered a lower quality allograft. Predictive models could learn these patterns and propagate existing disparities. |
| Post-deployment considerations | Potentially unbiased AI tools may lead to biased outcomes when deployed in real life either due to differential conditional distribution of outcomes of interests across subpopulations. | - Computer vision has been shown to aid detection of traditional adenomas; however, there are limited data on proximal and sessile serrated lesions which are more prevalent in black individuals. |

HCC, hepatocellular carcinoma; MELD, Model for End-Stage Liver Disease.
of patients—is two to four times more common in men. Therefore, the inclusion of this variable in the model inadvertently deprivatises women and perpetuates sex disparities in liver transplantation: women are up to 20% less likely to receive a liver transplant and 8.6% more likely to die on the transplant waitlist.

While conventional prediction models use few variables that appear to be transparent, high-capacity ML algorithms may employ innumerable variables from large volumes of data and identify highly complex non-linear patterns that are less comprehensible—that is, black box models—with the promise of increased predictive accuracy. Regardless of the type of model used, the variables selected (theme 3) as inputs for these models—conventional and AI-based alike—may appear objective at face value but can unwittingly introduce bias and lead to inequitable outcomes as illustrated with the MELD.

Bias due to variable selection is intrinsically related to the introduction of bias during algorithm development. Datasets used for predictive modelling may have unintended encoded biases, which has the potential to generate biased algorithms in the algorithm development phase (theme 4) as ML models aim to fit the datasets on which they train. For example, review of OPTN registry data reveals that medically underserved groups are less likely to be referred for liver transplant, less likely to undergo liver transplantation and more likely to receive lower quality allografts compared with white patients. Predictive models trained on such datasets could recreate these biases and amplify existing racial and ethnic liver transplantation disparities when deployed. Assigning transplant priorities based on predicted outcomes from biased models has huge ramifications for health equity in organ allocation. Identifying and rectifying biases after the model has been deployed can prove to be difficult—it took several years to show that an estimated glomerular filtration rate equation widely used to assign renal transplant priorities was biased against black patients. Predictive models trained on such datasets could recreate these biases and amplify existing racial and ethnic liver transplantation disparities when deployed. Assigning transplant priorities based on predicted outcomes from biased models has huge ramifications for health equity in organ allocation. Identifying and rectifying biases after the model has been deployed can prove to be difficult—it took several years to show that an estimated glomerular filtration rate equation widely used to assign renal transplant priorities was biased against black patients. Therefore, it is imperative that fairness and potential biases are addressed upfront.

Colorectal cancer screening

Colorectal cancer (CRC) prevention and control are major public health contributions by gastroenterologists. Effective CRC screening depends significantly on the endoscopist’s ability to identify and remove high-risk colon and rectal polyps during colonoscopy. Adenoma detection rate (ADR) is a validated measure of colonoscopy quality and significant predictor of adverse outcomes when the datasets used to develop models do not reflect the racial, ethnic, age, sex and gender diversity of our society to maximise generalisability in routine practice. Historically, research studies have not been conducted in settings that regularly serve these populations resulting in their ongoing exclusion. Therefore, it is critical to consider where marginalised populations are being.

Recent advances in clinical practice

In increasing equity in AI: potential solutions

It is imperative that we identify and implement pragmatic solutions that emphasise and optimise health equity in AI/ML development and application in gastroenterology and hepatology. Tools are needed to debias data collection, model training, model outputs and clinical application. The recently increased focus on equity in healthcare has motivated discussion about how to achieve these goals; these approaches are also urgently relevant to our field. Potential solutions to the equity challenges we have highlighted in this piece include incorporating a health equity lens early and often in AI/ML research and development, increasing the diversity of patients involved in AI/ML clinical trials, regulatory standards for reporting, and pre-deployment and post-deployment auditing (table 2).

First, a health equity approach to AI/ML requires technically diverse research teams that are aware of how bias can creep into all aspects of the research continuum. Beyond this, gastroenterology and hepatology research teams that employ AI/ML methods should engage health equity experts early in their work so that potential sources of bias are identified early and are addressed in a robust and effective manner.

Second, it is vitally important to increase the diversity of patient populations who are involved in algorithm development and validation in gastroenterology and hepatology. Data collection in AI in our field is currently limited by overfitting and spectrum bias. Overfitting occurs when models are closely tailored to a training set, which can reduce overall generalisability of the model when other datasets are used. Spectrum bias occurs when the datasets used to develop models do not reflect the diversity of the population they are meant to serve. Data-sets used for AI in gastroenterology and hepatology are often collected via retrospective or case-control design which poses risk for spectrum bias. Ideally, all algorithms should be developed and tested using a population that reflects the racial, ethnic, age, sex and gender diversity of our society to maximise generalisability in routine practice. Historically, research studies have not been conducted in settings that regularly serve these populations resulting in their ongoing exclusion. Therefore, it is critical to consider where marginalised populations are being.
served and how best to reach them, both in AI and non-AI contexts. Partnerships between gastroenterology practices, clinics and health centres that provide care for these populations can be leveraged to extend reach and promote generalisability when conducting research studies to advance equity. In addition, non-traditional settings where vulnerable populations routinely receive services should be considered to diversify representation in AI/ML studies and ensure equity in algorithm performance. Furthermore, models should be externally validated with new patient populations and datasets to limit the potential for spectrum bias and overfitting.81

Beyond diversifying the training data, it is crucial that labels (or data classifiers) used in prediction models are adequately representative of the desired outcome alone and are independent of societal inequities.11 It is also important to carefully consider the different conditional distributions of labels across subgroups and any variations in how they are classified and measured—these may suggest a need for optimising benchmarks or developing separate models for different subgroups.83 Some tools like Datasheets for Datasets14 or Model Cards for Model Reporting82 do exist, but identifying the precise cause of bias can be challenging and requires careful audits by multidisciplinary teams.

A third focus should be on regulatory standards. Mandating explicit reporting of descriptive data of the patient populations used in AI/ML development—such as race, ethnicity, income, insurance and sex—is a necessary step, as long as privacy is protected. Doing so enables a clear assessment of appropriate representation in the algorithm’s training dataset and the generalisability of its results. This type of descriptive data will also provide insight regarding which algorithms and models may not represent certain patient groups adequately.

Fourth, there must be robust processes in the pre-deployment phase to audit model outputs and ensure equal algorithmic performance for diverse patient populations. Sensitivity analyses evaluating algorithmic performance in subgroups can identify biased models with inequitable outcomes. Pre-emptive efforts to adjust models before deployment and mass dissemination protect marginalised subpopulations from inequitable outcomes and can also have cost-saving implications—the excess cost of racial health disparities in the USA is estimated at $230 billion over a 4-year period.84 For effective impact, the definition of ‘inequitable outcome’ set by regulators must be fair, clear, specific and quantifiable. Ongoing surveillance in the post-deployment setting is also imperative to monitor for unintended consequences of AI/ML and confirm unbiased algorithmic performance in actuality. Of note, access to training data and prediction methodologies of most large-scale AI/ML algorithms is frequently restricted, thus limiting independent efforts to assess for algorithmic biases and how they may have arisen.11 This reality underscores the importance of deidentified open-access data sharing in accordance with FAIR87 data principles—findability, accessibility, interoperability and reusability—which could be highly instrumental in promoting health equity by providing insight into which AI/ML algorithms could perpetuate and/or exacerbate disparities.

Finally, combining AI/ML models with physician clinical decision-making—that is, an augmented intelligence approach with a physician-in-the-loop configuration—may be beneficial in generating ethical and equitable AI/ML tools.88 Augmented intelligence may be of bidirectional utility as AI/ML models can standardise approaches where considerable provider variability exists while physician interaction can help limit biases that may arise from these tools. However, this approach must be done with careful consideration as biases can also arise from physician interaction with prediction models including automation bias (over-reliance on prediction models), feedback loops, dismissal bias (conscious or unconscious desensitisation) and allocation discrepancy.89

While these efforts can minimise bias and create more ethical AI tools, they do not serve as substitutes for repairing medical mistrust50 91 and certainly do not obviate the structural changes needed to build a more equitable health system.92–94

CONCLUSIONS
We describe five themes to illustrate how AI/ML can lead to inequities in gastroenterology/hepatology, examples of the impact on health equity and several potential actionable solutions to ensure equity in AI. By the year 2045, white individuals will comprise less than 50% of the US population, thus this work is critical as AI/ML becomes more common globally and the USA becomes more diverse.92

Our primary limitation was the inability to measure or quantify inequities in each clinical example provided. Though each example provided relates directly to a major theme of mechanisms of inequities in AI/ML, the degree to which each specific example led to bias cannot be directly measured. In addition, we did not have access to all of the model information used to develop the algorithms discussed nor to robust cost information that could enable a review of cost implications of current AI approaches. This fact highlights the importance of transparency to enable researchers’ access to data and inputs included in each algorithm to advance equity. Lastly, the examples provided in this paper are not an exhaustive list but rather focus on strong and relevant illustrations of how prediction models and AI/ML algorithms in gastroenterology and hepatology can lead to biased systems and inequitable health outcomes.

Table 2 Approaches to eliminate bias in AI/ML

<table>
<thead>
<tr>
<th>Approaches to eliminate bias in AI/ML</th>
<th>Description</th>
</tr>
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<tbody>
<tr>
<td>Appropriate research expertise</td>
<td>Involve health equity experts in the conception, development and deployment of AI/ML</td>
</tr>
<tr>
<td>Diverse study populations</td>
<td>Diversify study populations to adequately represent marginalised populations in training datasets. Convenience samples, such as datasets from electronic health records, claims data and so on, may not be adequately representative of marginalised groups.</td>
</tr>
<tr>
<td>Diverse study settings</td>
<td>Expand research locations to non-conventional settings where traditionally under-represented and vulnerable populations can be easily reached such as community health centres, faith-based organisations, barbershops, community service organisations and other settings.</td>
</tr>
<tr>
<td>Regulatory measures</td>
<td>Determine fair, clear, specific and quantifiable regulatory measures of inequitable outcomes. Researchers should be required to report descriptive data on study populations by sex, race, ethnicity as long as privacy is protected. Standards should be consistent across regulatory bodies, peer-reviewed scientific journals and gastroenterology/hepatology professional societies.</td>
</tr>
<tr>
<td>Pre-deployment auditing</td>
<td>Mandate auditing processes and sensitivity analyses to assess algorithmic performance across subpopulations in the pre-deployment phases.</td>
</tr>
<tr>
<td>Post-deployment auditing</td>
<td>Establish auditing processes to assess algorithmic performance across subpopulations in the post-deployment phase and pathways for rapidly mitigating bias if discovered in the post-deployment phase.</td>
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AI, artificial intelligence; ML, machine learning.
There are several key strengths of this paper. First, we provide clear and actionable solutions to address health equity in AI/ML that can be used by researchers and clinicians alike. Second, we provide concise themes that illustrate how AI/ML can lead to health inequity in gastroenterology matched to specific examples. Our overarching goal is to increase attention to an important potential downside of AI as its use becomes more prevalent and pervasive in the fields of gastroenterology and hepatology.

Here, we adapt a framework to consider equity in AI/ML algorithms used in gastroenterology/hepatology and a platform for discussion around an increasingly relevant topic. In other fields of medicine, we have started to reassess prediction models and algorithms and incorporate a health equity lens. The field of gastroenterology and hepatology has already taken a leading role in clinical applications for AI in medicine, and it is therefore especially important that, as a field, we take a leading role in ensuring that equity considerations are emphasized. This framework will help gastroenterology/hepatology researchers and clinicians prioritize equity in AI/ML development, implementation, and evaluation so that we can give every patient an opportunity to benefit from the technological advances that the future brings.

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