LLMs and Their Applications in Medical Artificial Intelligence

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Medical artificial intelligence (AI) is a cross-disciplinary field focused on developing advanced computing and AI technologies to benefit medicine and healthcare. Globally, medical AI has tremendous potential to support the United Nations' sustainable development goals pertaining to health and well-being. In particular, large language models (LLMs) afford opportunities for positively disrupting medical AI-related research and practice. We present a research framework for LLMs in medical AI. Our framework considers the interplay between health and well-being goals, disease lifecycle stages, and the important emerging role of LLMs in medical AI processes related to various lifecycle stages. As part of our framework, we describe the LLM multiplex - important multimodal, multi-model, multicultural, and multi-responsibility considerations for LLMs in medical AI. We discuss how the five articles in the special issue relate to this framework and are helping us learn about the opportunities and challenges for LLMs in medical AI.

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1 Introduction

The United Nations (UN) sustainable development goals (SDGs) include a set of goals related to health and well-being. Globally, these health-related SDGs have motivated a number of public health initiatives and nation-specific long-term targets. For instance, in the United States, Public Health 3.0 calls for timely, actionable, and granular intelligence to improve health and health equity of communities [8, 15]. In Europe, there is discussion on how to advance the UN's health and well-being SDGs across the European Union [11]. "Japan 2035" details the nation's public health targets and aspirational state for their national health system [26]. Similarly, "Healthy China 2030" outlines the country's vision for public health services, the medical industry, as well as food and drug safety [29]. In all such public health initiatives, medical artificial intelligence (AI) has an important role to play.

Within the medical AI space, large language models (LLMs) present tremendous opportunities for research and practice that enhances health and well-being outcomes at scale. Accordingly, the purpose of this article and accompanying special issue is to foster a dialogue on a research agenda

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for LLMs in medical AI. We present a research framework for LLMs in medical AI that considers the interplay between health and well-being goals, disease lifecycle stages, and the important emerging role of LLMs in medical AI processes related to various lifecycle stages. As part of our framework, we describe the LLM multiplex - important multimodal, multi-model, multicultural, and multi-responsibility considerations for LLMs in medical AI. We also describe how the five articles in the special issue relate to this framework and are elucidating opportunities and challenges for LLMs in medical AI.

2 A Research Framework for LLMs in Medical AI

Figure 1 presents our research framework for LLMs in medical AI. The framework is comprised of four components: (1) relevant health and well-being goals; (2) the four disease lifecycle stages; (3) different types of medical AI processes; and (4) the LLM multiplex. Details are as follows.

2.1 Health and Well-being Goals and Disease Lifecycles

As depicted at the top of Figure 1, the UN's health and well-being SDG includes reducing infant and child mortality rates, maternal mortality, diseases, enhancing mental health and wellness, reducing substance abuse, as well as other important health considerations [22]. A classic mantra in medical research and practice is "prevention is better than cure." This saying refers to the four stages of the disease lifecycle (depicted in the second part of Figure 1): prevention, diagnosis, treatment, and management [27]. Prevention involves periodic checkups, health lifestyle and diet practices, as well as routine screenings [8]. Diagnosis entails identifying illnesses and diseases as well as providing accurate patient prognoses. Treatment relates to alleviation of illness and disease through interventions such as medication, therapy, surgery, etc. [27]. The management stage entails recovery, rehabilitation, and related post-treatment activities. The lifecycle stages are the states and progressions by which the illnesses and diseases associated with the aforementioned well-being goals manifest and progress in individual patients. In the ensuing subsection, we describe how medical AI processes informed by the LLM multiplex can enhance lifecycle outcomes (bottom part of Figure 1).

2.2 Medical AI Processes and the LLM Multiplex

Two common ways in which AI can enhance existing processes is via AI-enabled automation and AI-embedded augmentation [1]. The former entails replacing manual human processes such as diagnosis, question answering, or chat help/support with AI-enabled diagnoses (i.e., machine learning predictions), question answering (LLMs for Q&A), or chat bots. Additionally, with the rise of generative AI, machine learning models such as LLMs can perform an increasing array of zero or few-shot assessment/scoring use cases, as well as generative AI use cases such as generating multimedia responses, having conversations, producing knowledge graphs, and so on [2]. For medical AI processes, this creates a 2x2 of AI-in-the-loop combinations in terms of automation/augmentation and assessment/generation (depicted in the third section of Figure 1).

The state-of-the-art for natural language processing has seen a few major shifts in recent years in text representation learning. Namely, from static word embeddings to contextual embeddings to transformer-based language models culminating with the rise of LLM foundation models [2, 4, 5, 20]. When envisioning the role of LLMs in medical AI, particularly as part of a global public health narrative, a key consideration is the LLM multiplex. The LLM multiplex is a set of multi-dimensional LLM-related factors that are salient in an array of contexts, including medical AI processes for any and all disease lifecycle stages. Depicted in the bottom part of Figure 1, these factors include: multimodal, multi-model, multicultural, and multi-responsibility. LLMs and Their Applications in Medical Artificial Intelligence

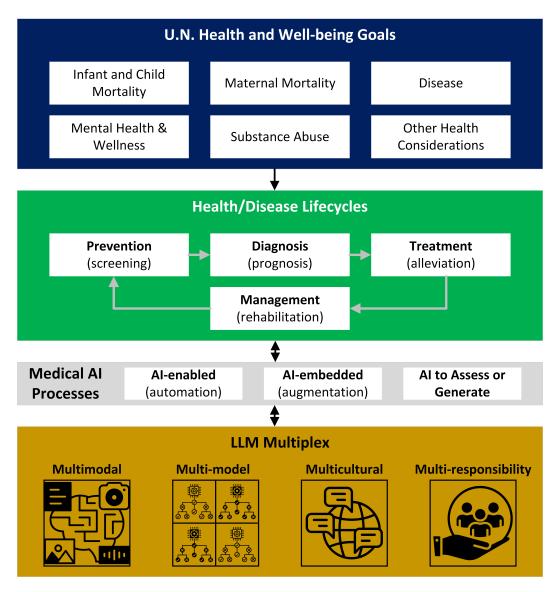


Fig. 1. A framework for LLMs in Medical AI

Multimodal refers to the fact that for any medical AI processes encompassing LLMs, patient data is inherently multimodal, potentially spanning speech text, audio, visual, and spatio-temporal input signals [13, 25]. *Multi-model* pertains to the use of modeling pipelines, architectures, and medical AI systems involving multiple models running in parallel, sequentially, and/or arranged in elaborate ensembles [13]. *Multi-cultural* relates to the importance of considering cultural artifacts such as language, customs, beliefs, values, and norms [16, 32]. Recent studies have noted a paucity of research on cultural alignment in LLMs [3]. *Multi-responsibility* relates to the importance of the various tenets of responsible AI, including fairness, privacy, transparency (i.e., explainability and interpretability), security, and the role of the human in the loop [2, 14]. The fairness considerations

in medical AI include disparate impact and lack of equal opportunities [1, 12, 24, 31]. LLMs can potentially exacerbate these concerns across protected attributes such as various demographic dimensions [10, 18], both for upstream representational harm and downstream allocational harm [17]. Similarly, transparency considerations such as interpretability and explainability of diagnosis, prognosis predictions, and optimal treatment strategies are crucial for garnering physician and patient buy-in [28]. Moreover, whereas LLM privacy and security is a major concern in various contexts [6, 19], the sensitive nature of medical AI makes them an issue of paramount importance. We believe this LLM multiplex - the multimodal, multi-model, multi-cultural, multi-responsibility environments where LLMs have great potential to inform and enhance global public health - are a crucial aspect of the research framework for LLMs in medical AI. In the following section, we describe how the five articles in the special issue shed light on important facets of this framework.

2.3 Articles in the Special Issue

Figure 2 depicts the five articles in the special issue (rows in the figure) and how they relate to the research framework (columns in the figure). Liu et al. [21] examine federated learning strategies for preserving privacy collaboratively fine-tuning LLMs. They focus on the important LLM tasks such as question answering and analyzing patient-physician medical dialogue using two medical testbeds (as well as datasets from other domains). Their work has implications for various health and well-being goals related to the prevention and diagnosis lifecycle stages, with implications for AI-enabled generation. In regards to the LLM multiplex, their work aligns with multi-responsibility - how to automatically generate answers to patient questions while preserving privacy of those on which the LLM was fine-tuned.

Qin and Cook et al. [23] review and benchmark LLMs for detecting depression from speech to text transcriptions of remote patient-physician interviews. Their work is closely aligned with the mental health global crisis that is overburdening psychiatrists and related mental health professionals. They focus on diagnosis - how well LLMs can detect depression from remote patient interviews. Their research relates to AI-enabled assessment - using LLMs for fine-tuned classification. They connect with the LLM multiplex facet of multimodal (by focusing on speech to text transcriptions). They also relate to multi-model by benchmarking the interaction effects between multiple automated speech recognition (ASR) models and various downstream classification LLMs.

Dai et al. [7] tackle visual question and answering by proposing a multimodal LLM that can answer questions involving analysis of text and images. Their research relates to various health and wellbeing goals mostly at the prevention and diagnosis stages, focusing on AI-enabled assessment and generation. They relate to the LLM multiplex aspect of multimodal LLMs that support biomedical vision-language understanding for automatic answering of user questions.

Wu et al. [30] evaluate the efficacy of building Chinese medical knowledge graphs using LLMs such as GPT-3.5 and GPT-4. Beginning with expert annotated knowledge, they use LLMs to construct the graphs. Their work has implications for various health and well-being goals across the prevention, diagnosis, and treatment stages. The work is a nice example of AI-embedded generation where the LLM augments the human experts' annotation efforts with graph construction support. It relates to the multi-cultural aspect of the LLM multiplex as multi-lingual knowledge of Chinese and English must be employed to effectively construct the graphs.

Dou et al. [9] design and develop a Chinese medication guidance system that can automate or augment medication guidance during the treatment phase. They employ a robust evaluation that includes domain experts and physicians, to ensure the LLM guidance is valid, complete, and contains appropriate warnings about adverse drug reactions. Their work relates to the LLM multiplex dimension of multi-cultural LLMs - to capture expert knowledge from local physicians and pharmacists, their work employs bilingual conversational LLMs. LLMs and Their Applications in Medical Artificial Intelligence

Paper	Health & Well- being Goals	Lifecycle Stage(s)	Medical Al Processes	LLM Multiplex
Differentially Private Low-Rank Adaptation of Large Language Model Using Federated Learning [21]	Question answering for various health & well-being goals	Prevention, Diagnosis	Al-enabled generation: Automatic patient question answering	Multi-responsibility: Ensure privacy through federated learning
Language Models for Online Depression Detection: A Review and Benchmark Analysis on Remote Interviews [23]	Mental health and wellness	Diagnosis	AI-enabled assessment: Automated depression detection	Multimodal and Multi-model: Assess patients' speech to text transcriptions for multiple ASR-LLM combinations
MOSS-MED: Medical Multimodal Model Serving Medical Image Analysis [7]	Visual question answering for various health & well-being goals	Prevention, Diagnosis	Al-enabled assessment & generation: Automatic user question answering for medical image analysis	Multimodal: Support biomedical vision-language understanding
Zero-Shot Construction of Chinese Medical Knowledge Graph with GPT-3.5-turbo and GPT-4 [30]	Knowledge graph construction for various health & well-being goals	Prevention, Diagnosis, Treatment	Al-embedded generation: Human expert annotation coupled with LLMs for graph construction	Multi-cultural: Build a knowledge graph for the Chinese medical domain using GPT
ShennongMGS: An LLM-based Chinese Medication Guidance System [9]	Medication guidance for various health & well-being goals	Treatment	Al-embedded & Al-enabled generation: Human experts/physicians evaluate the LLM outputs in regards to advice validity, completeness, etc.	Multi-cultural: Build a Chinese medication guidance system using bilingual conversant LLMs

Fig. 2. How Articles in the Special Issue Relate to the Framework

3 Conclusions

We conclude with a few possible research directions we anticipate will become more pervasive in the coming years as the research on LLMs in medical AI becomes increasingly robust:

- *From AI Processes to Full-stack AI Companions* Presently, most LLM artifacts proposed for medical AI focus on one or two stages of the disease lifecycle. As personalized LLMs "meet" precision medicine, research that explores the use of LLMs across all stages of the lifecycle, acting as personalized AI companions to patients, will be at a premium.
- A Multiplicity in the LLM Multiplex The current LLM medical AI research landscape is exploring one or two facets of the LLM multiplex. As LLMs get further integrated into medical AI processes, research is needed to design fully fleshed out multiplex solutions encompassing multiple modalities, models, cultures, and responsible AI tenets.
- Complex Medical AI Processes Spanning Automation/Augmentation and Assessment/Generation - Most existing research is focusing on medical AI processes with a single automation or augmentation and assessment or generation characteristic. Consistent with the shift towards omni-lifecycle, LLM multiplex intensive research and practice, we anticipate a need for research on increasingly sophisticated medical AI processes that integrate various combinations of the medical AI process 2x2.

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