Evidence Retrieval and Grounding in Medicine

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Agent

Aid Kit



Department of Electrical and Computer Engineering Boston University

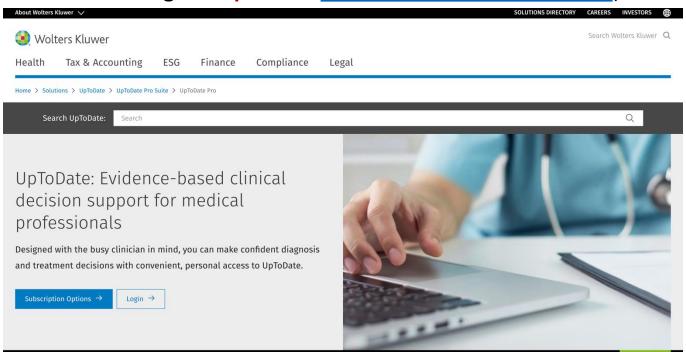
Outline

- Background
- Related Works and Motivation
- MedPodGPT
- PodGPT
- Agentic System



Part 1 – Background

■ Traditional search engine: UpToDate (\$6,713 million revenue, 2024)



Credit: https://www.uptodate.com/contents/search

Why Up	ToDate?	Product	Editorial	Subscription Options
All	Adult	Pediatric	Patient	Graphics
Showin	a rosults fo	r what is the	rocommon	ded treatment of chronic incomnia in adults

Overview of the treatment of insomnia in adults

... three active treatment arms (CBT-I, zolpidem, or both) in 63 young and middle-aged adults with chronic insomnia, there were no differences in total sleep time among active treatment groups and... every six months is recommended. In patients who fail insomnia treatment, it is important to discuss expectations of sleep, particularly for older adults and those with comorbidities....

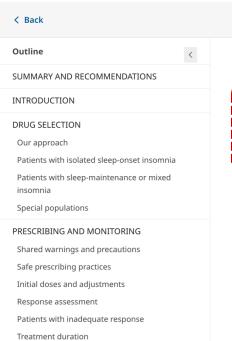
Pharmacotherapy for insomnia in adults

... the preferred first-line treatment for chronic insomnia in adults and has been endorsed as first-line therapy by multiple societies and guideline panels Pharmacotherapy should not be the sole **treatment** for insomnia. CBT-I is **recommended** as first-line treatment for chronic insomnia

Cognitive behavioral therapy for insomnia in adults

... This approach is a **recommended treatment** for insomnia disorder and is a multistep process carried out over multiple sessions. During the treatment,... and delivery of CBT-I and other behavioral treatments for insomnia in adults. An overview of the treatment of insomnia and pharmacologic therapies for insomnia are presented separately.... behavioral therapy for insomnia (CBT-I) is a multicomponent treatment for chronic insomnia disorder that aims to identify and target the multiple...

Credit: https://www.uptodate.com/contents/search



Pharmacotherapy for insomnia in adults



INTRODUCTION

Insomnia is one of the most common symptoms for which adults seek medical advice. While pharmacotherapy should not be the sole treatment for insomnia, medication may be part of an integrated approach that includes behavioral strategies and treatment of relevant comorbidities.

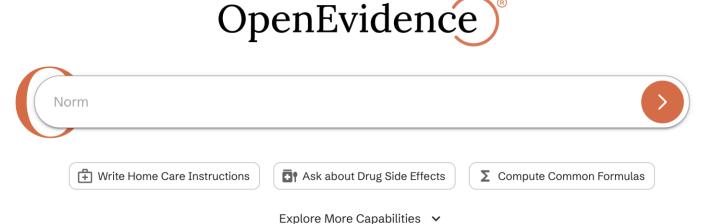
A range of medications and substances are used to treat insomnia in adults, spanning the following categories:

- Medications with regulatory approval for treatment of insomnia
- Off-label prescription medications prescribed for insomnia
- Over-the-counter (OTC) sleep aids (eq, diphenhydramine and doxylamine, alone or in "PM" formulated medications)

Credit: https://www.uptodate.com/contents/search

Part 1 – Background

Al-powered search engine: OpenEvidence (\$1 billion valuation, 2025)



Credit: https://www.openevidence.com/

Partnering with OpenEvidence: A Life-Saving Healthcare Revolution

This AI-powered search engine for doctors could be one of the most important companies of the next decade.

What is the latest diagnosis of dementia?

→ Finished thinking

The latest diagnostic criteria for dementia are based on a combination of clinical assessment and, increasingly, biomarker evidence, with criteria evolving to reflect advances in the understanding of neurodegenerative diseases.

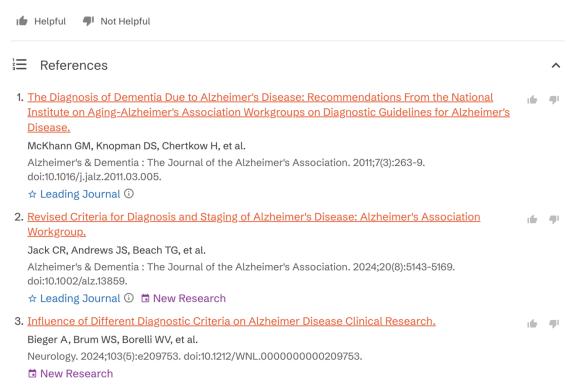
For all-cause dementia, the National Institute on Aging and the Alzheimer's Association (NIA-AA) recommend that dementia is diagnosed when there is evidence of cognitive or behavioral (neuropsychiatric) symptoms that interfere significantly with the ability to function at work or in usual daily activities. This determination is made through clinical judgment, based on history from the patient and a knowledgeable informant, and supported by cognitive testing. The diagnosis requires impairment in at least two cognitive domains (such as memory, executive function, language, visuospatial skills, or personality/behavior), and the deficits must represent a decline from previous levels of functioning and not be explained by delirium or major psychiatric disorder. The NIA-AA emphasizes that the distinction between mild cognitive impairment and dementia is based on the degree of functional impairment, with dementia requiring significant interference in daily life. [1]

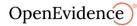
Recent updates from the Alzheimer's Association Workgroup further incorporate biomarkers into the diagnostic process, particularly for Alzheimer's disease. The 2024 criteria define Alzheimer's disease as a biological process, with diagnosis possible in asymptomatic individuals based on abnormal amyloid and tau biomarkers (e.g., amyloid PET, CSF, or plasma phosphorylated tau). An abnormal "Core 1" biomarker is sufficient to establish a diagnosis of Alzheimer's disease, while "Core 2" biomarkers (such as tau PET) provide prognostic information and increase diagnostic confidence. The criteria also recognize the importance of integrating clinical and biological staging, acknowledging that comorbidities and cognitive reserve may influence the clinical presentation. [2-3]

In summary, the most up-to-date consensus is that dementia is diagnosed clinically by evidence of acquired cognitive decline interfering with daily function, with increasing use of biomarkers for extinlesized specificity, especially in Algheimer's disease, as recommended by the National Institute



In summary, the most up-to-date consensus is that dementia is diagnosed clinically by evidence of acquired cognitive decline interfering with daily function, with increasing use of biomarkers for etiological specificity, especially in Alzheimer's disease, as recommended by the National Institute on Aging and the Alzheimer's Association.[2]

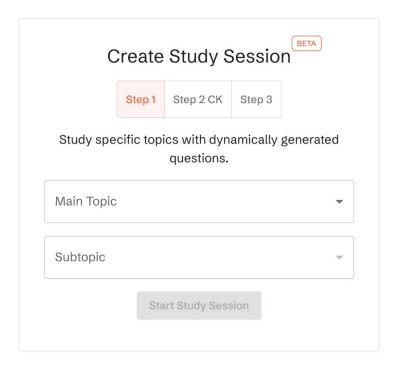








USMLE Test Prep



Study Session (i) View Session Stats

A 29-year-old female presents to the clinic with a chief complaint of fatigue and weight gain over the past six months. She works as a high school teacher in a rural area with limited access to healthcare specialists. The patient reports feeling cold all the time and has experienced constipation and mild depression. She also notes her periods have become irregular, and she has had difficulty concentrating at work. She denies any recent illnesses or significant stressors.

Her past medical history is significant for a head injury from a motor vehicle accident two years ago, which required hospitalization but did not result in any lasting neurological deficits. She takes no regular medications and denies smoking, alcohol, or illicit drug use.

On physical examination, the patient appears tired and has a dry, pale skin texture. Her vital signs are: blood pressure 110/70 mmHg, heart rate 58 bpm, temperature 96.8°F (36°C), and respiratory rate 14 breaths per minute. Thyroid examination reveals no palpable goiter. Reflexes are delayed, and she has mild facial puffiness with periorbital edema. No other abnormalities are noted.

Laboratory results reveal:

- TSH: 0.4 μIU/mL (normal: 0.5-4.5 μIU/mL)
- Free T4: 0.6 ng/dL (normal: 0.8-1.8 ng/dL)
- Free T3: 1.8 pg/mL (normal: 2.3-4.2 pg/mL)
- Serum cortisol (8 AM): 5 µg/dL (normal: 5-25 µg/dL)
- Prolactin: 25 ng/mL (normal: 4-23 ng/mL)

Additional imaging with MRI of the brain reveals a 1 cm lesion in the pituitary region suggestive of a pituitary adenoma.

A 29-year-old female presents to the clinic with fatigue, weight gain, cold intolerance, constipation, mild depression, irregular periods, and difficulty concentrating. She has a history of a head injury from a motor vehicle accident two years ago. Physical examination reveals dry, pale skin, delayed reflexes, mild facial puffiness, and periorbital edema. Laboratory results show low TSH, free T4, and free T3 levels, with a slightly elevated prolactin level. An MRI of the brain reveals a 1 cm lesion in the pituitary region suggestive of a pituitary adenoma. What is the most appropriate next step in management?

0	(A) Schedule the patient for surgical removal of the pituitary adenoma
0	(B) Order a fine-needle aspiration biopsy of the pituitary lesion
0	(C) Initiate levothyroxine therapy
0	(D) Refer to an endocrinologist for further evaluation and management
0	(E) Start dopamine agonist therapy

Credit: https://www.openevidence.com/

USMLE-style multiple-choice question answering:



Medical Students



Residents

Need harder questions:

Fellows Doctors (Clinicians/Physicians)



from a motor vehicle accident two years ago. Physical examination reveals dry, pale skin, delayed reflexes, mild facial puffiness, and periorbital edema. Laboratory results show low TSH, free T4, and free T3 levels, with a slightly elevated prolactin level. An MRI of the brain reveals a 1 cm lesion in the pituitary region suggestive of a pituitary adenoma. What is the most appropriate next step in management?

(A) Schedule the patient for surgical removal of the pituitary adenoma

(B) Order a fine-needle aspiration biopsy of the pituitary lesion

(C) Initiate levothyroxine therapy

 $\hfill \bigcirc$ (D) Refer to an endocrinologist for further evaluation and management

(E) Start dopamine agonist therapy

∠ Large
✓ Ask a follow-up question...

Submit





OpenEvidence

complications. [1-0]

Endocrinologists are uniquely qualified to distinguish between functioning and nonfunctioning adenomas, interpret laboratory artifacts such as the hook effect, and initiate appropriate first-line therapies (e.g., dopamine agonists for prolactinomas, surgical referral for nonfunctioning or other functioning adenomas). [2][6:9] The Endocrine Society and other expert guidelines recommend comprehensive endocrine evaluation in all patients with pituitary adenomas, including assessment for hormone hypersecretion and hypopituitarism, as well as formal visual field testing if there is concern for optic chiasm involvement. [1-2][4] Early specialist involvement is critical to prevent morbidity from untreated hormone deficiencies, guide further imaging or surgical referral if indicated, and optimize long-term outcomes. [1-2][4]

In summary, the complexity of diagnosis and management, the need for tailored hormone replacement, and the potential for significant morbidity underscore the necessity of endocrinology referral in this clinical scenario.

∃ References

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The Journal of Clinical Endocrinology and Metabolism. 2011;96(4):894-904. doi:10.1210/jc.2010-1048.

^

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- Jama. 2023;329(16):1386-1398. doi:10.1001/jama.2023.5444.
- ☆ Leading Journal
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4. <u>Hormonal Replacement in Hypopituitarism in Adults: An Endocrine Society Clinical Practice</u>
<u>Guideline.</u>

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The Journal of Clinical Endocrinology and Metabolism. 2023;108(9):2400-2423. doi:10.1210/clinem/dgad174.

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- ☆ Leading Journal

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- ☆ Leading Journal

Part 1 – Background

Patient background

A 27-year-old man presents to the emergency department. He was brought in by staff from the homeless shelter when they found him unresponsive. The patient is a known IV drug abuser but otherwise has an unknown past medical history. He currently attends a methadone clinic. His temperature is 99.5°F (37.5°C), blood pressure is 97/48 mmHg, pulse is 140/min, respirations are 29/min, and oxygen saturation is 98% on room air. Initial laboratory values are shown below. Serum: Na+: 139 mEq/L. Cl-: 100 mEq/L. K+: 6.3 mEq/L. HCO3-: 17 mEq/L. Glucose: 589 mg/dL. The patient is given treatment. After treatment, his temperature is 99.5°F (37.5°C), blood pressure is 117/78 mmHg, pulse is 100/min, respirations are 23/min, and oxygen saturation is 98% on room air. His laboratory values are seen below. Serum: Na+: 139 mEq/L. Cl-: 100 mEq/L. K+: 4.3 mEq/L. HCO3-: 19 mEq/L. Glucose: 90 mg/dL.

Part 1 – Background

- Question and options
 - Question

What is the best next step in management?

- Options
 - A. Insulin, potassium, IV fluids, and glucose
 - B. IV fluids only
 - C. Oral rehydration
 - D. Supportive therapy and close monitoring
- Answer
 - A. Insulin, potassium, IV fluids, and glucose

- (1) Multiple choice question-answering
- (2) Open-ended question-answering

Part 2 – Related Works

■ Question-answering without rationale → lack explainability

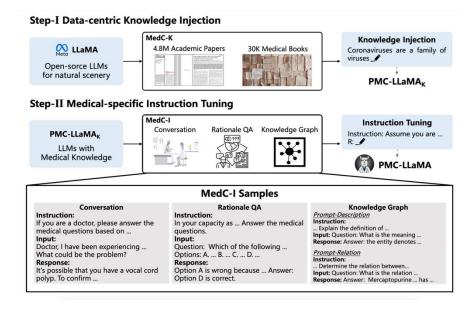
Dataset	Source		Description						
			Finetuning						
Medical Flash Cards	Anki Flas	shcards	Rephrased Q&A pairs derived from the front and back sides of medical flashcards						
Stack Exchange	Academia	a	Q&A pairs generated from questions and their top-rated answers	39,633					
	Biology			7,482					
Medical Flash Cards	Fitness			3,026					
	Health			1,428					
	Bioinforn	natics		906					
Wikidoc	Living Te	extbook	Q&A pairs generated from paragraphs, where questions were for- mulated from rephrased paragraph titles, and answers were ex- tracted from paragraph text						
	Patient tion	Informa-	Q&A pairs generated from paragraph headings and associated text content	5,942					
			Evaluation						
USMLE	Step 1		Multiple choice questions from the USMLE self-assessment with image-based questions excluded	119					
	Step 2			120					
	Step 3			135					

Instruction tuning

(Supervised fine-tuning)

Part 2 – Related Works

■ Question-answering with rationale → lack factuality



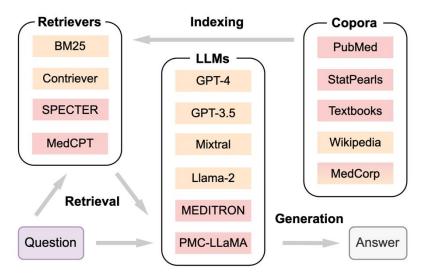
###Rationale: This patient's clinical presentation and laboratory approximatives suggest a diagnosis of thrombotic thrombocytopenic purpura (TTP). TTP is a microangiopathic hemolytic anemia, resulting from severe deficiency or functional abnormality of the cleaving protease ADAMTS13, leading to accumulation of ultra-large multimers of von Willebrand factor, platelet adhesion, and widespread thrombi in microcirculation. Clinically, TTP is characterized by the pentad of thrombocytopenia, microangiopathic hemolytic anemia, neurologic abnormalities, renal insufficiency, and fever. This pentad is not always complete, and high clinical suspicion should be maintained even if not all five elements are present, as in this case. In terms of management, plasma exchange therapy is the mainstay and should be initiated as soon as the diagnosis is suspected, even before laboratory confirmation, given the severity and rapid progression of TTP. This therapy helps by removing the autoantibodies that inhibit ADAMTS13 and replenishing the deficient enzyme through plasma replacement. Options A and B, high-dose glucocorticoids and cyclophosphamide and rituximab, respectively, can be used in addition to plasma exchange in certain refractory or relapsed cases but are not first-line treatment. Option C, Vancomycin and cefepime, are antibiotics used to treat infections, which doesn't align with this patient's presentation. ###Answer: OPTION D IS CORRECT.

Credit:

- [1] Wu et al., PMC-LLaMA: Towards Building Open-source Language Models for Medicine, In Journal of the American Medical Informatics Association'23.
- [2] Qiu et al., Towards Building Multilingual Language Model for Medicine, In Nature Communications' 24.

Part 2 – Related Works

- Question-answering with retrieval-augmented generation (RAG)
 - Bias problem ← no further evidence assessment
 - Low efficiency ← model doesn't decide whether to use evidence



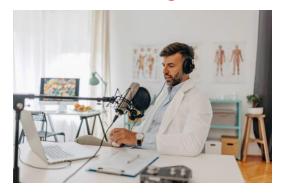
Part 2 – Motivation

- Question-answering without rationale
 - Clinical decision-making with rationale → explainability
- Question-answering with rationale
 - Clinical decision-making with external evidence → factuality
- Question-answering with RAG
 - Clinical decision-making with evidence assessment ← bias problem
 - Clinical decision-making with an automatic and dynamic process — low efficiency

Credit:

- [1] Han et al., MedAlpaca: An Open-Source Collection of Medical Conversational AI Models and Training Data, In arXiv'23.
- [2] Wu et al., PMC-LLaMA: Towards Building Open-source Language Models for Medicine, In Journal of the American Medical Informatics Association'23.
- [3] Qiu et al., Towards Building Multilingual Language Model for Medicine, In Nature Communications' 24.
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What: A multilingual audio-augmented LLM for medical research and education



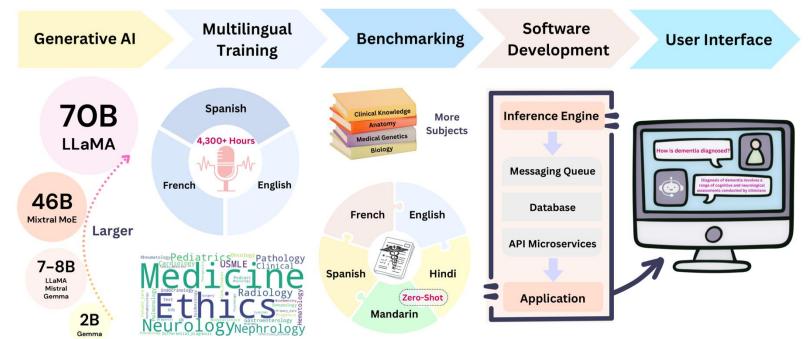


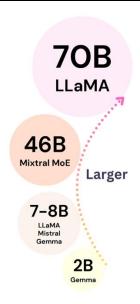




- Why: Up to date and domain-relevant information ← podcasts
- Where: Journals, exam preparation materials, and clinical practice
- When: Most recent (e.g., since 2023)
- How: Continual pretraining of large language models (LLMs)

A multilingual audio-augmented LLM for medical research and education





Large language models

- 2B Gemma
- 7B Gemma, 7B Mistral, 8B LLaMA
- 46B Mixtral 8×7B MoE
- 70B LLaMA

$$\mathcal{L}_{\pi_{\theta}} = -\sum_{i} \log(\pi_{\theta}(x_{i}|\mathbf{x}_{< i})).$$

 π_{θ} : LLM, parameterized by θ

 $\mathbf{x} = [x_1, x_2, ..., x_t]$: a sequence of texts







Podcasts



CURRENT ISSUE

✓ SPECIALTIES

✓ TOPICS

MULTIMEDIA

LEARNING/CME

AUTHOR CENTER

Q

Podcasts & Feeds

INTENTION TO TREAT NOT OTHERWISE SPECIFIED Incisive analysis of critical and Exploring some of health timely issues in medicine and care's toughest challenges and INTENTION TO TREAT health care. areas of greatest promise. Recent episodes. Recent episodes. FOLLOW: FOLLOW: **NEJM THIS WEEK NEJM INTERVIEWS** A summary of what's in this Current topics in medicine and health care, with authors week's issue. and editors. Recent episodes. Recent episodes.

FOLLOW:



THE NEJM AI GRAND ROUNDS PODCAST

Informal conversations with a variety of experts exploring the deep issues at the intersection of artificial intelligence, machine learning, and medicine.

Recent episodes.



FOLLOW: 🔞 🖨

- Podcasts (journals)
 - OpenAI Whisper automatic speech recognition (ASR) model (2022)
 - Audio → texts (transcripts)

Journal Podcasts											
Podcast	Language	Episodes	Audio Time	Mean Length Episode	Number of	Mean Text Tokens					
Toucast	Language	Episodes	(min)	$\pm\sigma$ (min)	Text Tokens	per Episode $\pm\sigma$					
NEJM	English	1974	39256.0	19.89 ± 9.74	4,760,783	1928.22 ± 14.87					
JAMA	English	2235	32163.0	14.39 ± 8.66	3,454,191	1928.64 ± 15.54					
The Lancet	English	2029	28279.0	13.94 ± 7.62	3,300,982	1925.89 ± 20.88					
The BMJ	English	300	13264.2	44.21 ± 10.35	2,235,458	1897.67 ± 75.07					
Annals Latest Highlights	English	396	6427.0	16.23 ± 8.09	803,958	1927.96 ± 14.60					
Annals On Call	English	142	3440.0	24.22 ± 3.65	522,547	1928.22 ± 15.64					
Pediatrics on Call	English	98	3299.0	33.66 ± 6.09	565,781	1930.99 ± 15.00					
Procedure Ready: Ob/Gyn	English	20	383.7	19.19 ± 5.00	63,667	1929.30 ± 13.41					
Revista Médica AFP Podcast	Spanish	40	1055.0	26.38 ± 3.70	190,518	1924.42 ± 16.34					

Credit:

[1] https://openai.com/index/whisper/

[2] Jia et al., MedPodGPT: A Multilingual Audio-augmented Large Language Model for Medical Research and Education, In medRxiv'24.

Podcasts (test preparations)

Test Preparation Podcasts											
Podcast	Languaga	Episodes	Audio Time	Mean Length of Episode	Number of	Mean Text Tokens					
Toucast	Language	Lpisoues	(min)	$\pm\sigma$ (min)	Text Tokens	per Episode $\pm\sigma$					
Divine Intervention Podcasts	English	480	18363.8	38.26 ± 24.07	2,269,153	1931.19 ± 13.53					
The Radiology Review Podcast	English	127	2517.7	19.82 ± 10.10	292,949	1927.29 ± 26.81					
Crush Step 1: The Ultimate USMLE Step 1 Review	English	49	2176.2	44.41 ± 15.31	328,194	1930.55 ± 13.03					
The USMLE Guys Podcast	English	31	1464.3	47.24 ± 43.47	156,923	1937.32 ± 6.12					
Harrison's PodClass: Internal Medicine Cases and Board Prep	Spanish	95	905.2	9.53 ± 2.24	101,574	1916.49 ± 22.77					
El Interno Desvelado	Spanish	4	99.13	24.78 ± 11.91	17,121	1902.33 ± 25.54					
Curso MIR Asturias	Spanish	3	17.7	5.89 ± 4.53	3,872	1936.00 ± 9.00					

Podcasts (clinical experts)

Clinical Podcasts											
Podcast	Language	Episodes	Audio Time	Mean Length of Episode	Number of	Mean Text Tokens					
roucast	Language	Lpisoues	(min)	$\pm \sigma$ (min)	Text Tokens	per Episode $\pm \sigma$					
The Curbsiders Internal Medicine Podcast	English	485	28749.2	59.39 ± 16.08	5,772,083	1929.82 ± 17.06					
This Podcast Will Kill You	English	168	18363.8	38.26 ± 24.07	2,269,153	1931.19 ± 13.53					
The Clinical Problem Solvers	English	315	13500.1	42.86 ± 14.51	2,493,777	1927.18 ± 23.32					
PsychEd: educational psychiatry podcast	English	62	3556.3	57.36 ± 17.52	607,237	1927.74 ± 16.36					
Run the List	English	97	1973.0	20.34 ± 6.44	352,977	1928.84 ± 15.17					
Goljan Pathology Lectures	English	37	1886.0	50.97 ± 4.58	412,086	1934.68 ± 13.45					
Core IM: 5 Pearls	English	54	1847.1	34.21 ± 10.19	361,213	1931.62 ± 10.16					
Neurology Clinical Pearls	English	27	333.2	12.34 ± 3.19	42,494	1931.54 ± 10.78					
Tutorías Medicina Interna	Spanish	570	19834.9	34.80 ± 25.01	4,311,263	1898.39 ± 64.31					
Leucocitos isotópicos	Spanish	68	2537.8	37.32 ± 9.55	481,676	1797.29 ± 154.42					
Medicina Con Cabeza	Spanish	246	2457.8	9.99 ± 3.44	462,383	1902.81 ± 57.55					
Medicina de impacto	Spanish	57	2406.5	42.22 ± 9.13	492,363	1915.81 ± 29.28					
Ronda, El Podcast de Medicina Interna	Spanish	20	1084.4	54.22 ± 25.01	206,218	1891.91 ± 71.90					
Medicina De Bolsillo Hablando de Medicina	Spanish	45	958.3	21.30 ± 10.79	186,268	1844.24 ± 124.82					
La Tertulia de Cajal	Spanish	27	876.3	32.46 ± 18.28	186,001	1897.97 ± 57.71					
PedCast: Dos Pediatras y un Podcast	Spanish	14	458.5	32.75 ± 10.62	89,127	1896.32 ± 58.05					
Neurobiologie et Immunite	French	21	1882.8	89.66 ± 14.77	383,189	1896.97 ± 40.12					
Incubateur Néonat	French	25	1579.3	63.17 ± 21.28	391,475	1918.99 ± 24.15					
Guideline.care	French	68	1369.1	20.13 ± 6.30	293,301	1917.0 ± 29.29					
La Minute Rhumato	French	119	921.0	7.74 ± 2.19	132,354	1918.17 ± 23.59					
Oncologie cellulaire et moléculaire - Hugues de Thé	French	11	852.9	77.53 ± 19.81	186,693	1905.03 ± 44.42					
Le podcast des Conférenciers (UFR3S) by Université de Lille	French	65	768.4	11.82 ± 19.58	86,105	1913.44 ± 42.78					
Super Docteur	French	47	676.3	14.39 ± 6.50	139,824	1915.40 ± 33.26					
Médecine, Sciences et Recherche clinique	French	24	332.2	13.84 ± 4.58	63,314	1918.61 ± 26.60					
NéphrOdio	French	40	318.6	7.96 ± 2.58	55,716	1921.24 ± 19.59					
La Minute Néonat	French	37	307.6	8.31 ± 1.93	57,435	1914.50 ± 31.58					
Le Med G Eclairé	French	11	249.2	22.66 ± 16.76	51,988	1925.48 ± 12.57					
La Minute du Pancréas	French	22	209.4	9.52 ± 2.34	38,376	1918.80 ± 23.34					
L'essentiel des principales pathologiesaå	French	14	151.3	10.81 ± 13.10	23,098	1924.83 ± 11.25					
AR-Pod le Podcast de lanesthésie-réanimation	French	12	139.0	11.59 ± 4.52	22,998	1916.50 ± 26.48					

Credit: Jia et al., MedPodGPT: A Multilingual Audio-augmented Large Language Model for Medical Research and Education, In medRxiv'24.

Benchmarks (in-domain performance)

		Model											
Language	Benchmark Datasets	Gemma	a 2B	Gemm	a 7B	Mistra	l 7B	LLaMA 3 8B		Mixtral MoE		LLaMA 3 70B	
		Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Ours
	MedExpQA	15.20	23.40	34.40	45.20	47.20	46.20	57.60	61.40	52.80	61.20	78.40	77.60
	MedMCQA	34.81	35.24	40.66	44.86	42.65	45.50	58.64	58.82	50.20	53.54	71.12	70.58
	MedQA	29.69	33.38	38.26	44.56	46.27	47.80	61.12	59.21	54.05	53.22	77.85	77.51
	PubMedQA	47.80	55.50	63.40	55.30	51.60	41.75	59.40	49.20	42.80	32.20	73.00	75.75
	Anatomy	43.70	42.04	49.63	52.96	56.30	56.67	68.89	69.82	64.44	68.15	77.04	77.78
English	Clinical Knowledge	41.51	38.78	55.47	62.17	61.89	62.26	72.08	73.68	67.92	74.90	82.26	83.40
	College Biology	44.44	47.05	61.11	68.06	61.81	64.93	74.31	77.43	72.92	77.95	91.67	92.36
	College Medicine	36.99	37.14	50.29	55.06	57.80	59.97	67.05	68.06	63.58	69.07	78.61	78.18
	Medical Genetics	43.00	44.75	54.00	66.00	64.00	65.50	80.00	77.25	70.00	78.00	91.00	91.00
	Professional Medicine	29.78	34.10	50.37	60.02	56.99	63.33	76.84	75.64	72.06	73.07	90.44	90.26
	Average	36.69	39.14	49.76	55.42	54.65	55.39	67.59	67.05	61.08	64.13	81.14	81.22
	FrenchMedMCQA	29.91	28.43	29.60	40.27	45.48	44.32	41.74	44.63	55.14	58.02	63.24	73.05
	MedExpQA	19.20	20.60	26.40	39.20	40.80	41.20	48.00	43.60	50.40	56.00	76.80	74.00
	Anatomy	35.56	35.18	48.15	49.63	33.33	39.45	45.19	47.41	55.56	59.63	67.41	68.52
	Clinical Knowledge	32.45	36.51	50.94	57.92	55.47	53.02	61.89	61.13	65.66	71.51	78.87	80.56
French	College Biology	33.33	38.02	46.53	52.78	53.47	49.65	57.64	62.50	67.36	72.92	86.81	87.67
	College Medicine	32.95	35.84	43.93	47.98	51.45	48.56	57.80	59.40	57.80	63.44	69.94	74.71
	Medical Genetics	35.00	40.00	50.00	57.25	47.00	59.00	66.00	67.00	71.00	72.00	90.00	89.50
	Professional Medicine	24.26	28.95	33.09	42.00	43.38	43.84	51.47	55.51	59.56	64.15	72.79	73.34
	Average	30.33	32.94	41.08	48.38	46.30	47.38	53.72	55.15	60.31	64.71	75.73	77.67
	HeadQA	33.77	34.32	48.21	54.47	53.79	55.54	59.66	61.24	64.77	68.00	81.44	82.44
	MedExpQA	21.60	23.00	32.80	38.40	46.40	40.40	40.00	43.00	52.80	52.40	73.60	76.60
	Anatomy	37.78	39.08	42.22	51.11	45.93	49.63	48.15	52.96	60.74	62.22	71.11	74.44
	Clinical Knowledge	37.74	38.78	53.96	55.47	54.34	56.13	58.49	62.08	68.68	68.40	78.49	80.00
Spanish	College Biology	29.17	35.94	48.61	50.35	55.56	56.25	54.86	55.04	66.67	69.10	85.42	84.20
~~	College Medicine	32.37	34.39	43.93	48.84	54.34	48.99	49.71	54.05	59.54	58.24	69.94	72.97
	Medical Genetics	32.00	34.75	46.00	59.50	53.00	57.25	72.00	68.00	67.00	66.75	86.00	86.75
	Professional Medicine	26.47	30.06	38.24	43.56	47.06	45.68	51.84	50.74	53.68	56.90	69.49	68.94
	Average	31.36	33.79	44.25	50.21	51.30	51.23	54.34	55.89	61.74	62.75	76.94	78.29

Benchmarks (zero-shot cross-lingual transfer performance)

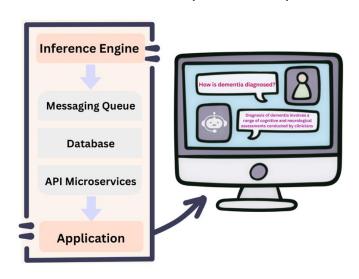
		Model											
Language	Benchmark Datasets	Gemma	a 2B	Gemma	a 7B	Mistra	l 7B	LLaMA	3 8B	Mixtral	MoE	LLaMA	3 70B
		Baseline	Ours										
	MedQA-MCMLE	33.39	33.43	40.51	45.98	39.67	39.25	63.63	66.32	45.80	47.14	84.68	83.73
	Anatomy	28.38	23.98	25.00	31.59	25.00	30.41	33.78	35.98	33.11	26.35	63.51	64.02
	Clinical Knowledge	29.11	28.27	31.22	39.87	33.33	32.60	49.37	50.95	39.24	38.61	71.73	71.94
	College Medicine	28.94	32.23	33.70	36.08	30.77	30.68	52.01	56.50	38.46	40.94	75.82	80.49
Chinese	Medical Genetics	32.39	32.39	43.75	45.17	38.64	42.33	43.18	44.60	45.45	45.88	61.36	57.53
	Medical Nutrition	33.79	35.69	40.69	44.66	42.07	37.24	53.10	50.00	49.66	51.90	66.21	68.28
	Traditional Chinese Medicine	27.57	28.52	31.35	36.35	24.86	28.52	43.24	39.46	30.27	30.94	66.49	67.98
	Virology	37.28	36.98	46.15	54.44	43.79	48.22	59.76	58.88	53.25	50.15	76.33	77.51
	Average	31.36	31.44	36.55	41.77	34.77	36.16	49.76	50.34	41.91	41.49	70.77	71.44
	Anatomy	25.93	32.22	34.07	36.86	23.70	30.00	40.00	35.18	31.11	34.44	52.59	57.78
	Clinical Knowledge	26.42	28.96	41.89	41.04	24.91	35.85	48.30	46.70	38.11	36.70	63.40	69.06
	College Biology	26.39	33.16	26.39	34.03	19.44	28.47	32.65	37.16	30.56	32.81	58.33	68.06
Hindi	College Medicine	24.86	27.60	42.20	43.35	23.12	33.09	41.04	43.64	27.17	33.24	60.69	64.74
	Medical Genetics	31.00	30.50	36.00	41.75	28.00	29.25	46.00	45.75	40.00	43.25	71.00	77.00
	Professional Medicine	25.37	26.19	30.88	41.08	22.06	28.67	36.40	39.34	29.41	29.50	45.59	64.70
	Average	26.66	29.77	35.24	39.69	23.54	30.89	40.73	41.29	32.73	34.99	58.60	66.89

Software development

- vLLM: inference engine
- Apache Cassandra: query optimization
- Flask: store chats and conversations in Cassandra and send text inference requests to a queue
- RabbitMQ: queue requests
- PostgreSQL: RAG database management
- OAuth 2.0: Authorization and user management

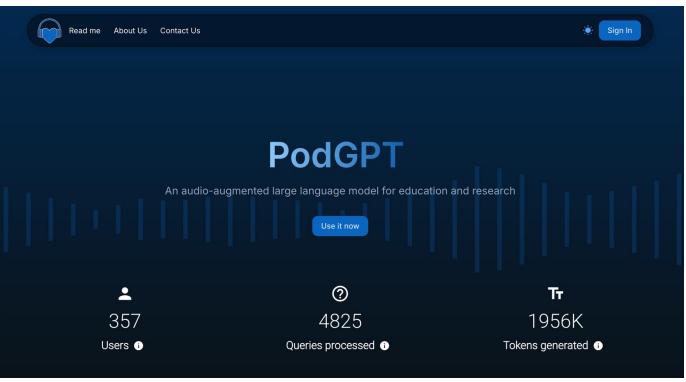
Credits:

- [1] PodGPT: https://podgpt.org/
- [2] vLLM: https://github.com/vllm-project/vllm
- [3] Apache Cassandra: https://github.com/apache/cassandra
- [4] Flask: https://github.com/pallets/flask
- [5] RabbitMQ: https://github.com/rabbitmq/rabbitmq-server
- [6] PostgreSQL: https://github.com/postgres/postgres
- [7] OAuth 2.0: https://github.com/postgres/postgres



Software development

https://podgpt.org





- 1. Customized teaching cases: "I'm teaching second-year medical students. Generate a case on a patient with a chronic obstructive pulmonary disease (COPD) exacerbation with diagnostic uncertainty about heart failure exacerbation."
- **2. Expert discussion**: "I have a medical case in 4 parts. As our diagnostic expert, provide the problem representation, a prioritized differential, and your choice of next test for each part."
- 3. Summarization and feedback:
- "Would my patient be included in this study?" or "Summarize methodological points of the study."
- "I need a 2-paragraph summary of these course evaluations. Include 2 direct quotes highlighting learner qualities."
- "I've written these patient instructions aiming for an 8th grade reading level. Determine their current reading level and offer feedback for improvement."

Credit: Rodman et al., Using Generative Artificial Intelligence in Medical Education, In Academic Medicine'25.

Take away

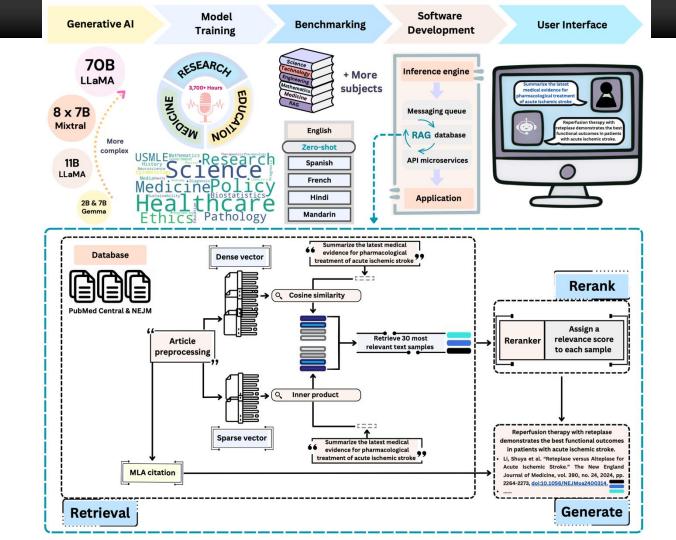
- Podcasts are valuable sources of up to date and domain-relevant information
- Continual pretraining of LLMs on podcasts can enhance overall model performance
- Improve in-domain (language) performance
- Improve zero-shot multilingual transfer capabilities

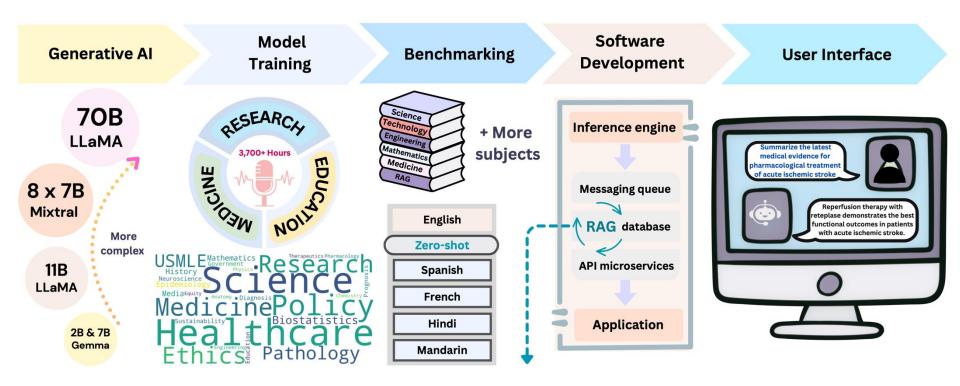
Part 4 – PodGPT

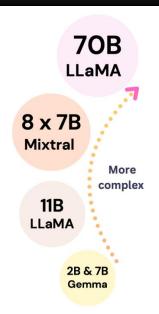
- What: An audio-augmented LLM for research and education
 - Fields: Science, technology, engineering, mathematics, and medicine (STEMM)



- Why: (1) Up to date information ← podcasts (2) Factual information ← journal articles
- Where: Creative Commons Attribution (CC BY) podcasts from journals/experts, and NEJM
- When: Most recent (e.g., since 2023)
- How: Continual pretraining of LLMs







Part 4 – PodGPT

- Large language models
 - 2B Gemma
 - 7B Gemma, 7B Mistral
 - Quantized 70B LLaMA (11B)
 - 46B Mixtral 8×7B MoE
 - 70B LLaMA







Part 4 – PodGPT

Podcasts

Podcast	Episodes	Audio time (min)	Mean length episode $\pm \sigma$ (min)	Number of text tokens	Mean text tokens per episode $\pm \sigma$
NEJM This Week	457	13,300.17	29.10 ± 2.92	2,029,219	4440.30 ± 419.35
NEJM Interviews	654	9,223.44	14.10 ± 8.15	1,732,879	2649.66 ± 1522.35
NEJM Core IM Internal Medicine Podcast	170	5,285.72	31.09 ± 10.08	1,077,154	6336.20 ± 2093.18
NEJM Curbside Consults	74	1,977.46	26.72 ± 11.04	408,189	5516.07 ± 2522.46
NEJM Clinical Conversations	108	1,829.66	16.94 ± 4.13	320,968	2971.93 ± 774.89
NEJM Leadership Conversations	100	1,765.76	17.66 ± 4.38	306,490	3064.90 ± 759.09
NEJM AI Grand Rounds	24	1,459.11	60.80 ± 14.84	303,499	12645.79 ± 3107.35
NEJM Intention to Treat	40	997.22	24.93 ± 5.27	169,632	4240.80 ± 954.41
NEJM Not Otherwise Specified	20	836.03	41.80 ± 15.71	146,718	7335.90 ± 2880.36
TWiV: This Week In Virology	1,186	104,089.57	87.77 ± 30.30	20,188,268	17022.15 ± 5785.86
TWiP: This Week In Parasitism	245	21,129.03	86.24 ± 15.35	4,381,511	17883.72 ± 3814.75
TWiM: This Week in Microbiology	320	20,641.50	64.50 ± 10.36	3,667,519	11461.00 ± 2121.78
TWiEVO: This Week In Evolution	100	8,845.09	88.45 ± 10.99	1,756,480	17564.80 ± 2517.80
IMMUNE	93	6,918.84	74.40 ± 19.67	1,363,176	14657.81 ± 3888.47
TWiN: This Week In Neuroscience	53	3,581.13	67.57 ± 10.40	644,712	12164.38 ± 2078.49
Matters Microbial	62	3,418.33	55.13 ± 11.33	637,647	10284.63 ± 2357.42
Infectious Disease Puscast	65	2,298.71	35.36 ± 5.96	415,145	6386.85 ± 1126.15
Urban Agriculture	29	2,171.63	74.88 ± 17.58	443,859	15305.48 ± 4214.82
On The Wards: On The Pods Medical Podcast for Doctors	245	5,915.14	24.14 ± 8.00	1,175,307	4797.17 ± 1781.74
Digital Campus Podcast	64	3,169.44	49.52 ± 5.85	605,977	9322.72 ± 1528.99
emDOCs.net Emergency Medicine (EM) Podcast	112	1,576.72	14.08 ± 4.30	303,672	2711.35 ± 864.29
Policy in Plainer English Podcast	73	1,089.20	14.92 ± 7.52	208,580	2857.26 ± 1509.80
Open Minds from Creative Commons	21	803.28	38.25 ± 12.79	145,624	6619.27 ± 2722.90
What is Global Health?	18	486.47	27.03 ± 10.02	87,653	4869.61 ± 2054.00
Consilience Sustainability In Progress (SIP) Podcast	9	403.95	44.88 ± 17.89	70,461	7829.00 ± 3535.75
Research Pulse: Future Focussed Health Insights	16	177.79	11.11 ± 2.29	33,672	2104.50 ± 476.85
Our People: Central to Healthcare	9	161.15	17.91 ± 7.42	31,545	3505.00 ± 1474.57

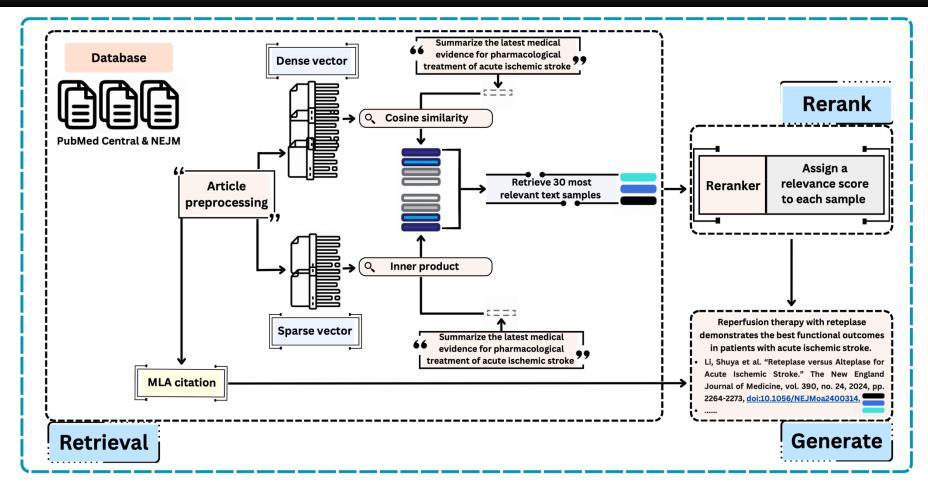
Credit: Jia et al., PodGPT: An Audio-augmented Large Language Model for Research and Education, In Nature npj Biomedical Innovations'25.

Part 4 – PodGPT

Benchmarks (in-domain performance)

Model	Ger	Gemma 2B		Gemma 7B		Quantized LLaMA 70B		Mixtral 8×7B MoE		. 70B
MMLU benchmark	Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Ours
Physics	29.85	30.74 (0.12)	42.38	47.64 (0.64)	76.79	79.24	56.08	56.19 (0.71)	78.63	79.26
Biology	44.82	46.58 (0.45)	63.32	66.43 (0.58)	90.97	91.51	72.14	77.84 (0.90)	93.53	93.53
Chemistry	30.04	30.96 (0.62)	43.42	44.14 (1.31)	69.94	73.15	46.60	50.33 (1.34)	70.16	69.66
Computer Science	40.59	44.20 (0.11)	53.58	54.62 (0.29)	80.24	82.30	59.25	60.98 (1.25)	78.72	79.44
Engineering	39.31	42.07 (0.49)	44.14	47.94 (0.60)	73.79	73.79	57.24	56.38 (0.90)	75.17	75.17
Mathematics	25.69	26.44 (0.57)	34.97	39.23 (0.16)	68.83	71.34	47.42	46.59 (1.01)	63.97	64.18
Medicine	40.62	41.72 (0.15)	55.22	59.50 (0.14)	86.11	87.38	67.38	74.00 (0.71)	88.65	88.65
Average	35.85	37.53	48.15	51.36	78.10	79.82	58.02	60.33	78.40	78.56

						<u></u>	Model					
	Language	Benchmark datasets	Gen	nma 2B	Ger	nma 7B	Quanti LLaMA		Mixtral	8×7В МоЕ	LLaMA	70B
			Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Ours
		MedQA-MCMLE	33.54	33.14 (0.13)	40.81	45.20 (0.06)	82.69	84.15	45.94	45.49 (0.31)	86.14	86.02
		Physics	32.75	31.73 (0.37)	35.37	40.13 (0.54)	61.08	60.53	42.72	38.90 (1.42)	62.42	61.92
		Biology	24.26	25.44 (0.00)	33.14	37.43 (0.64)	62.72	63.91	44.38	39.94 (2.15)	59.76	60.36
		Chemistry	25.00	27.08 (0.32)	29.55	37.12 (0.54)	45.45	43.94	29.55	34.47 (0.85)	46.97	47.73
Benchmarks	Mandarin	Computer Science	32.24	34.18 (0.32)	40.78	47.92 (0.59)	73.60	76.82	57.16	56.20 (2.42)	78.46	78.46
- Delicillians		Engineering	33.40	32.05 (0.40)	41.72	45.58 (0.20)	62.48	63.72	42.30	46.75 (0.93)	66.92	66.16
		Mathematics	26.82	24.64 (0.65)	27.96	30.73 (0.81)	53.72	51.93	38.21	30.41 (0.90)	58.89	57.90
		Medicine	31.18	31.62 (0.15)	35.72	39.65 (0.18)	70.06	71.21	41.06	41.77 (0.74)	73.42	73.29
(zero-shot		Average	29.90	29.98	35.63	40.47	63.98	64.53	42.66	41.74	66.62	66.48
(-3:3 3::33		MedExpQA	19.20	22.40 (0.00)	28.00	37.40 (1.18)	78.40	77.60	49.60	52.80 (1.50)	77.60	77.60
		FrenchMedMCQA	31.46	28.04 (0.22)	33.64	43.69 (0.34)	81.62	84.11	57.01	64.64 (0.47)	87.23	87.85
transfer)		Physics	26.93	28.88 (0.23)	35.58	42.65 (0.59)	71.28	70.31	55.56	56.51 (0.51)	72.40	72.29
uansierj		Biology	34.51	39.35 (0.39)	50.16	57.04 (0.34)	89.29	91.01	70.03	73.52 (1.01)	90.90	90.90
	F	Chemistry	26.07	29.09 (0.32)	37.98	40.20 (0.53)	64.94	64.93	47.14	49.73 (0.45)	65.18	64.43
	French	Computer Science	35.46	37.40 (0.69)	44.20	45.23 (0.70)	75.85	75.88	57.54	56.43 (0.41)	76.49	76.24
		Engineering	38.62	36.20 (0.60)	46.90	47.59 (0.00)	68.28	72.41	53.79	54.31 (0.57)	71.72	71.72
		Mathematics	26.14	28.01 (0.51)	30.62	32.92 (0.98)	64.87	63.90	44.65	46.68 (1.14)	65.63	65.92
		Medicine	32.56	35.48 (0.12)	45.80	51.55 (0.18)	81.36	83.44	65.08	66.49 (0.29)	84.21	84.29
		Average	30.11	31.65	39.21	44.25	75.10	75.95	55.60	57.90	76.82	76.80
		Physics	25.49	26.60 (0.56)	29.32	33.39 (0.46)	58.48	55.90	32.06	31.14 (1.26)	58.68	59.12
		Biology	29.02	32.58 (0.18)	29.28	39.08 (0.66)	66.55	66.46	35.69	34.03 (0.74)	72.74	72.61
		Chemistry	24.08	20.88 (0.10)	35.26	36.97 (0.90)	50.84	51.05	33.98	30.25 (2.59)	54.04	53.29
	Hindi	Computer Science	32.15	30.30 (0.40)	36.64	41.49 (0.37)	65.78	61.22	37.20	36.40 (2.60)	66.45	66.45
	Hindi	Engineering	43.45	42.42 (0.34)	40.00	41.72 (1.04)	59.31	57.93	40.00	43.28 (1.23)	57.24	58.62
		Mathematics	25.33	24.87 (0.24)	29.33	30.96 (0.50)	53.19	48.35	33.74	33.10 (0.47)	54.60	52.82
		Medicine	26.77	29.07 (0.15)	34.00	40.26 (0.21)	64.60	65.08	33.98	33.43 (0.80)	71.04	71.05
		Average	29.47	29.53	33.40	37.70	59.82	58.00	35.24	34.52	62.11	61.99
		HEAD-QA	33.66	34.38 (0.10)	48.32	52.87 (0.19)	81.58	82.97	64.48	66.76 (0.50)	84.28	84.21
		MedExpQA	21.60	23.20 (0.00)	32.80	37.40 (0.35)	76.80	80.00	51.20	54.00 (1.65)	76.80	78.40
		Physics	28.06	28.86 (0.26)	40.64	43.63 (0.11)	70.20	70.92	53.32	54.84 (0.67)	71.59	72.13
		Biology	30.63	39.50 (0.55)	52.19	57.26 (0.51)	85.37	85.46	66.46	71.11 (0.47)	86.73	86.50
	Coonigh	Chemistry	27.06	25.94 (0.22)	35.98	39.93 (0.36)	63.72	61.72	48.36	48.74 (1.00)	67.18	65.93
	Spanish	Computer Science	37.09	40.43 (0.40)	45.93	48.04 (0.25)	73.38	74.30	56.74	56.71 (0.78)	74.77	75.72
		Engineering	43.45	38.79 (0.75)	47.59	49.14 (0.89)	73.10	73.10	55.17	53.79 (1.89)	72.41	71.72
		Mathematics	26.63	25.80 (0.14)	31.14	32.79 (0.39)	64.80	61.97	42.04	41.70 (0.72)	67.08	66.47
		Medicine	31.89	35.54 (0.15)	45.94	51.81 (0.44)	78.28	79.64	61.17	63.25 (0.90)	80.19	80.09
		Average	31.12	32.49	42.28	45.87	74.14	74.45	55.44	56.77	75.67	75.69



Credit: Jia et al., PodGPT: An Audio-augmented Large Language Model for Research and Education, In Nature npj Biomedical Innovations'25.

Part 4 – PodGP1

RAG Database

	JAMA Network Open	9,367	119,672	389.61
	The New England Journal of Medicine	2,013	33,344	400.33
	Cell	497	24,977	417.86
_	British Medical Journal	601	10,307	399.27
	The Lancet	458	9,425	403.77
	The Lancet Global Health	539	9,003	399.56
	Neurology	428	6,862	404.43
	JAMA Health Forum	524	5,524	390.70
	The Lancet Regional Health - Europe	337	5,435	398.94
	The Lancet Infectious Diseases	272	4,720	400.31
	The Lancet Regional Health – Western Pacific	243	3,267	397.28
	The Lancet Public Health Home	179	3,059	401.88
	JAMA Psychiatry	180	2,859	399.22
	JAMA Neurology	141	2,462	397.50
	The Lancet Oncology	116	2,304	400.62
	The Lancet Regional Health – Americas	174	2,262	395.02
	The Lancet Microbe	111	2,054	403.17
	The Lancet Psychiatry	119	1,951	400.77
	The Lancet Neurology	110	1,841	396.53
	JAMA Oncology	138	1,810	394.91
	The Lancet HIV	97	1,786	401.54
	The Lancet Planetary Health	100	1,776	400.43
	JAMA Pediatrics	135	1,773	391.99
	JAMA Internal Medicine	128	1,766	395.25
	The Lancet Regional Health – Southeast Asia	134	1,607	397.18
	The Lancet Respiratory Medicine	72	1,360	404.70
	The Lancet Child & Adolescent Health	68	1,205	402.95
	The Lancet Diabetes & Endocrinology	64	1,082	399.89
	The Lancet Gastroenterology and Hepatology	54	1,007	397.67
	JAMA Cardiology	60	947	396.69
	JAMA Surgery	68	941	399.31
	The Lancet Healthy Longevity	49	855	403.10
	JAMA Ophthalmology	56	810	398.01
	The Lancet Rheumatology	35	713	403.76
	The Lancet Haematology	33	604	398.36
	JAMA Dermatology	30	413	399.73
	JAMA Otolaryngology – Head & Neck Surgery	17	244	399.27

Journal

JAMA Facial Plastic Surgery

Number of articles | Number of text samples | Average length per sample

10

424.80

Part 4 – PodGPT

Benchmarks (in-domain performance)

	Benchmark d	atasets	MedExpQA	MedMCQA	MedQA	PubMedQA	MMLU Medicine	Average
		Baseline	19.20	34.71	29.54	46.80	40.62	34.17
	Gemma 2B	Ours	21.20 (0.69)	34.62 (0.02)	32.91 (0.15)	54.25 (0.54)	41.72 (0.15)	36.94
	Gemma 2B	Baseline+RAG	23.20	35.91	32.60	49.00	41.47	36.44
		Ours+RAG	28.00 (0.00)	35.96 (0.07)	34.43 (0.12)	51.95 (0.78)	42.12 (0.13)	38.49
		Baseline	34.40	40.69	37.78	61.80	55.22	45.98
	Gemma 7B	Ours	42.00 (0.89)	44.64 (0.09)	44.14 (0.21)	57.35 (1.37)	59.50 (0.14)	49.53
	Gennia /B	Baseline+RAG	35.20	40.64	39.28	61.40	54.14	46.13
		Ours+RAG	47.40 (1.18)	43.54 (0.07)	43.32 (0.25)	55.70 (1.88)	60.11 (0.39)	50.01
		Baseline	73.60	68.30	76.67	77.60	86.11	76.46
Model	Quantized LLaMA 70B	Ours	75.20	69.11	78.00	77.60	87.38	77.46
Model		Baseline+RAG	69.60	66.51	66.77	76.80	83.91	72.72
		Ours+RAG	76.00	68.13	69.60	78.80	85.62	75.63
		Baseline	52.80	50.11	54.20	42.00	67.38	53.30
	Mixtral	Ours	58.60 (1.54)	51.94 (0.31)	55.36 (0.81)	25.35 (0.78)	74.00 (0.71)	53.05
	8×7B MoE	Baseline+RAG	49.60	48.58	47.84	34.60	65.45	49.21
		Ours+RAG	57.12 (1.21)	51.29 (0.28)	51.14 (0.01)	24.44 (1.11)	72.01 (0.02)	51.20
		Baseline	75.20	70.31	78.40	76.20	88.65	77.75
	LLaMA 70B	Ours	75.20	70.62	78.32	76.40	88.65	77.84
	LLawiA 70B	Baseline+RAG	68.00	68.11	66.93	77.40	84.24	72.94
		Ours+RAG	71.20	68.59	68.42	77.40	85.60	74.24

Credit: Jia et al., PodGPT: An Audio-augmented Large Language Model for Research and Education, In Nature npj Biomedical Innovations'25.



Summarize the latest medical evidence for the effectiveness of crinecerfont in adult congenital adrenal hyperplasia.

Crinecerfont therapy allowed substantial and clinically meaningful glucocorticoid reduction to more physiological doses in adults with classic congenital adrenal hyperplasia and was associated with reduced adrenal androgen production. The treatment appeared to be acceptably tolerated, with similar incidences of treatment-emergent adverse events in both the crinecerfont and placebo groups, and achieved androstenedione control at week 24 in 74.6% of crinecerfont-treated participants compared with 52.6% with placebo.

 Auchus, Richard J., et al. "Phase 3 Trial of Crinecerfont in Adult Congenital Adrenal Hyperplasia." The New England Journal of Medicine, vol. 391, no. 6, 2024, pp. 504-514. doi:10.1056/NEJMoa2404656.



What is the association between gestational diabetes mellitus and cardiovascular disease?

Women with a history of gestational diabetes mellitus are at substantially higher risk of future cardiovascular and cerebrovascular diseases overall, with a pooled risk ratio of 1.72. This excess risk cannot be solely attributed to conventional cardiovascular risk factors, and is partially mediated by subsequent diabetes, highlighting the need for early intervention and continuous monitoring in women with gestational diabetes mellitus.

- Xie, Wenhui, et al. "Association of Gestational Diabetes Mellitus with Overall and Type Specific Cardiovascular and Cerebrovascular Diseases: Systematic Review and Meta-Analysis." BMJ, vol. 378, 21 Sep. 2022, e070244. doi:10.1136/bmj-2022-070244
- Echouffo Tcheugui, Justin B., et al. "Association of Concomitant Gestational Hypertensive Disorders and Gestational Diabetes With Cardiovascular Disease." JAMA Network Open, vol. 5, no. 11, 1 Nov. 2022, e22 (Score: 0.49 doi:10.1001/jamanetworkopen.2022.43618.



Summarize the latest medical evidence for monoclonal antibody L9LS to prevent malaria.

The monoclonal antibody L9LS has shown to be effective in preventing malaria, with 88% of participants protected after controlled human malaria infection, and protection was seen at serum concentrations as low as 9.2 µg per milliliter. A single low dose of L9LS administered subcutaneously may be a feasible strategy to limit malaria-associated morbidity and mortality among infants and young children in regions where malaria is endemic.

- Kayentao, Kassoum, et al. "Subcutaneous Administration of a Monoclonal Antibody to Prevent Malaria." The New England Journal of Medicine, vol. 390, no. 17, 2024, pp. 1549-1559. doi:10.1056/NEJMoa2312775. Score: 0.52
- Wu, Richard L., et al. "Low-Dose Subcutaneous or Intravenous Monoclonal Antibody to Prevent Malaria." The New England Journal of Medicine, vol. 387, no. 5, 2022, pp. 397-407. doi:10.1056/NEJMoa2 Score: 0.60



Explain the role of GPC3-Unc5 receptors.

GPC3-Unc5 receptors play a crucial role in regulating collective migration and correct targeting, and their signaling must be finely balanced to achieve effective migration. The interaction between GPC3 and Unc5 receptors elicits a repulsive cellular response, and disrupting or enhancing this interaction can impact various cellular processes, including apoptosis, migration, and survival, as seen in embryonal tumors and neuroblastoma samples.

• Akkermans, Onno, et al. "GPC3-Unc5 Receptor Complex Structure and Role in Cell Migration." Cell, vol. 185, no. 21, 2022, pp. 3931-3949.e26, doi:10.1016/j.cell.2022.09.025.



Are there disparities in household air pollution exposure between urban and rural areas?

There are disparities in household air pollution exposure between urban and rural areas, with exposures due to polluting fuels generally being much higher in rural settings than in urban settings. The average measured 24-hour HAP-PM2-5 personal exposure for users of biomass fuel with traditional stoves is 130 μ g/m³ in rural settings and 80 μ g/m³ in urban settings, indicating a significant difference in exposure levels between the two settings.

 Mohajeri, Nahid, et al. "Urban-Rural Disparity in Global Estimation of PM2.5 Household Air Pollution and Its Attributable Health Burden." The Lancet. Planetary Health, vol. 7, no. 8, 2023, pp. e660-e672, doi:10.1016/52542-5196(23)00133-X.
 Score: 0.55



What is the role of amygdala neurons in social decision making in primates?

The amygdala neurons in primates play a crucial role in social decision making by encoding object-specific values learned from social observation and own experience, and translating these values into representations of the partner monkey's forthcoming choices. This is evident from the fact that amygdala damage profoundly impairs primates' social behavior, and that distinct "simulation neurons" in the amygdala dynamically translate values into representations of the partner monkey's choices, supporting understanding of others' mental states.

 Grabenhorst, Fabian et al. "Primate Amygdala Neurons Simulate Decision Processes of Social Partners." Cell vol. 177,4 (2019): 986-998.e15. doi:10.1016/j.cell.2019.02.042

Part 4 – PodGPT

Performance on conversational dataset

Model		Gen	nma 2B			Gemma 7B		Quantized LLaMA 70B		Mixtral 8×7B MoE		LLaMA 70B	
		Baseline	Ours	Baseline	Ours	Baseline	Ours	Baseline	Baseline Ours		Ours		
Dornlovity	Original transcripts	17.54	10.15 (0.01)	27.04	8.75 (0.02)	7.34	7.27	6.11	6.32 (0.04)	7.31	7.10		
Perplexity	Our transcripts	14.08	7.64 (0.02)	22.72	6.67 (0.02)	6.12	5.51	5.23	5.05 (0.01)	6.10	5.55		

$$PPL(\mathbf{x}) = e^{-\frac{1}{t}\sum_{i}^{t} \log(p_{\theta}(x_{i}|\mathbf{x}_{< i}))}.$$

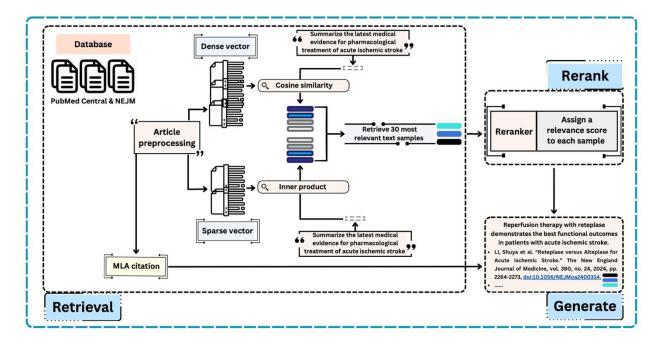
Part 4 – PodGPT

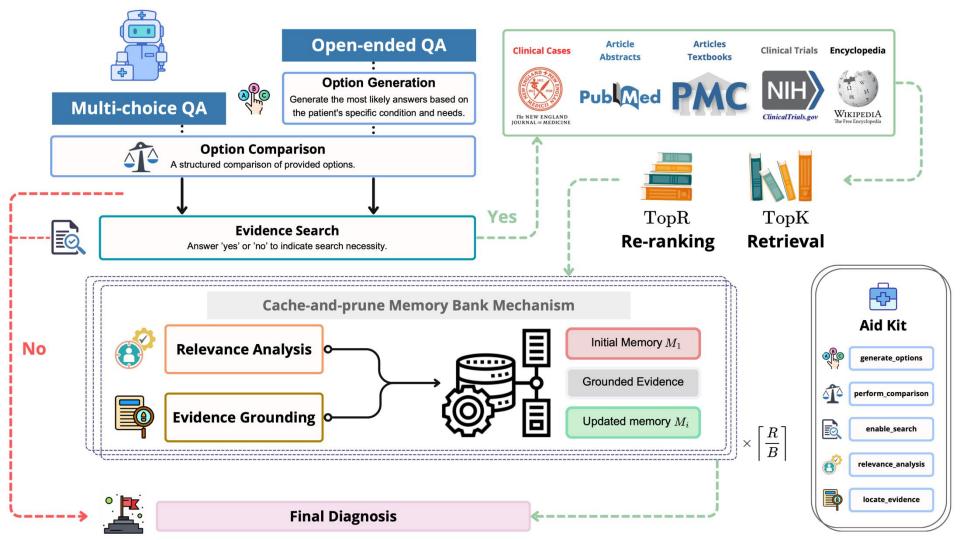
- Take away
 - Podcasts are valuable resources for STEMM research and education
 - Grounding LLMs with RAG enhances factual accuracy and reliability
 - Continual pretraining of LLMs on podcasts improves their conversational capability
 - Medical journals are valuable resources for evidence-based AI generation

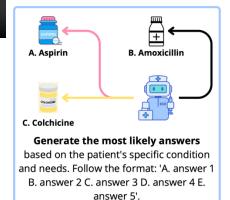
- Agentic memory-augmented retrieval and evidence grounding in medicine
 - Retrieval
 - Reranking
 - Generation
- Unified
- Automatic
- Dynamic

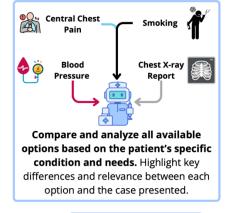














Indicate whether external evidence should be searched to support the patient's diagnosis based on the current clinical context and the need for additional information.



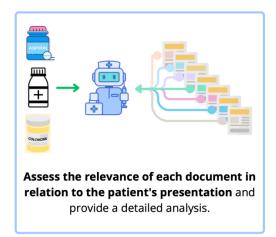
enable_search

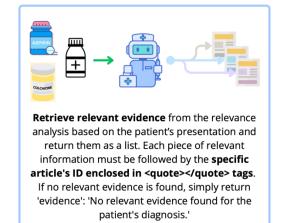


generate_options



perform_comparison







locate_evidence



relevance_analysis

1: Initialize Document Retriever ϕ , Evidence Reranker ψ

2: Initialize AI Agent π , Conversation C, Memory Bank M_1 3: Initialize Evidence database V

Algorithm 1 Agentic memory-augmented retrieval and evidence grounding system

4: Given patient background and question Q, instructions I, tools T5: All Agent π generates initial response $\prod_t \pi(y_t \mid T, Q, I, \mathbf{y}_{< t})$

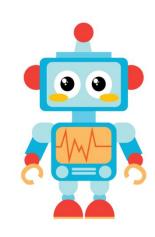
5: AI Agent
$$\pi$$
 generates initial response $\prod_t \pi (y_t \mid T, Q, I, \mathbf{y}_{< t})$

- 6: **while** tool calling **do**
 - Retrieve content from the tool calling to update conversation C7: 8:
 - if tool calling is enable_search then 9: Retrieve TopK documents arg TopK_{$\mathbf{v} \in \mathcal{V}$} $-\|\phi(\mathbf{x}) - \phi(\mathbf{v})\|_2$
 - Rerank TopR documents arg TopR_{$\mathbf{k} \in \mathcal{K}$} cos $(\psi(\mathbf{x}), \psi(\mathbf{k}))$
- 10: while $i \leq \lceil R/B \rceil$ do 11:
- Retrieve \mathcal{B}_i (a batch of \mathcal{R}) to update conversation C12: if tool calling is locate_evidence then 13:
- if Relevant document is grounded within <quote></quote> tags then 14: Update memory bank $M_i = \text{Prune}(M_{i-1} \cup \mathcal{B}_i)$
- 15: 16: end if
- end if 17:
- Remove \mathcal{B}_i from conversation C18:
- 19: end while until Sufficient information is gathered end if
- 20:
- 21: end while
- 22: if M_i then

26: **end if**

- **return** Final diagnosis $\prod_t \pi(y_t \mid T, Q, I, C, M_i, \mathbf{y}_{< t})$ 23: 24: **else**
- **return** Final diagnosis $\prod_{t} \pi (y_t \mid T, Q, I, C, \mathbf{y}_{< t})$ 25:

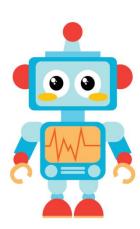
- Tool using (functional call)
- Enhanced interpretability and user trust
 - Perform a specific task with specific output constraints (but with input perturbations).
 - Better information retrieval with reduced hallucinations.
- Access to external tools, databases, and APIs
 - **Knowledge Acquisition** ← External knowledge
- Automation and efficiency
 - Complete complex tasks without excessive prompting.
- Interaction enhancement
 - Interact with external models, e.g., MRI model, CT scan model, etc.



- Tool using (functional call)
- Task planning
 - Which step should I solve?
- Tool selection
 - Should I use a tool(s)?
 - Which tool(s) should I use to solve this step problem?

Tool calling

- Calling and performing the specific action that the tool has defined.
- Response generation
 - Generating a response based on the tool's output description.



Tools

Tool Name	Parameter	Parameter Description
generate_options	answers	The most likely answers based on the patient's specific condition and needs.
perform_comparison	comparisons	A structured comparison of all options, detailing their relevance to the patient's case.
enable_search	search	Answer 'yes' or 'no' to indicate search necessity.
relevance_analysis	analysis	A comprehensive analysis detailing the relevance of each document to the patient's presentation, highlighting key matches, inconsistencies, and important findings.
locate_evidence	evidence	Relevant evidence applicable to the patient's presentation, with article IDs in <quote></quote> tags.

Database

Corpus	Number of Docs	Number of Snippets	Average Length	
PubMed Abstracts	23,897,881	23,897,881	290.01	
Wikipedia	6,458,670	29,642,311	166.47	
Clinical Trials	156,887	4,177,121	268.33	
PubMed Central Articles	123,194	8,155,929	202.46	
Textbooks	8,226	2,224,013	207.95	
Clinical Cases	1,479	17,821	215.61	

	Journal Title	Article Count	Journal Title	Article Count
	BMJ Open	37,488	JAMA Ophthalmol	434
	Proc Natl Acad Sci U S A	16,619	Lancet HIV	387
	JAMA Netw Open	10,824	BMJ Health Care Inform	366
- Dotoboso	Nature	8,148	JAMA Surg	287
Database	Cell	4,811	BMJ Neurol Open	282
	Science	4,660	JAMA Dermatol	279
	\mathbf{BMJ}	3,636	Lancet Psychiatry	270
	BMJ Glob Health	3,460	Lancet Public Health	264
	N Engl J Med	2,159	BMJ Support Palliat Care	262
	BMJ Open Qual	1,569	BMJ Nutr Prev Health	254
	JAMA	1,552	Lancet Respir Med	252
	BMJ Open Diabetes Res Care	1,434	JAMA Cardiol	239
	Lancet	1,344	Lancet Diabetes Endocrinol	225
	Neurology	1,216	Lancet Microbe	167
	BMJ Open Sport Exerc Med	1,201	BMJ Ment Health	167
	Lancet Reg Health West Pac	1,196	JAMA Otolaryngol Head Neck Surg	164
	BMJ Case Rep	1,190	Lancet Planet Health	162
	BMJ Paediatr Open	1,145	Lancet Haematol	157
	Lancet Reg Health Eur	1,077	BMJ Med	154
	BMJ Open Respir Res	1,031	Lancet Child Adolesc Health	154
	Lancet Reg Health Am	901	BMJ Evid Based Med	136
	Ann Intern Med	881	Lancet Digit Health	124
	Lancet Glob Health	805	BMJ Surg Interv Health Technol	120
	JAMA Intern Med	797	Lancet Gastroenterol Hepatol	117
	Lancet Infect Dis	676	BMJ Oncol	114
	BMJ Open Ophthalmol	656	Lancet Healthy Longev	102
	JAMA Neurol	639	BMJ Sex Reprod Health	100
	JAMA Health Forum	628	BMJ Mil Health	64
	BMJ Open Gastroenterol	625	Lancet Rheumatol	61
	Lancet Oncol	613	BMJ Open Sci	49
	BMJ Qual Saf	601	BMJ Innov	46
	JAMA Psychiatry	597	BMJ Simul Technol Enhanc Learn	42
	JAMA Pediatr	569	JAMA Facial Plast Surg	39
	BMJ Qual Improv Rep	547	Ann Intern Med Clin Cases	6
	JAMA Oncol	490	BMJ Outcomes	1
	Lancet Reg Health Southeast Asia	464	BMJ Clin Evid	1
	BMJ Public Health	453		
	Lancet Neurol	444	Total Number of Articles	123,194

Benchmarks (medical analysis and diagnosis)

_	Benchmark	Number of Testing Cases	Number of Choices	Cottings
!	USMLE Step 1 (40)	94	9	Getting
i	USMLE Step 2 (40)	109	6	more
Ų,	USMLE Step 3 (40)	122	6	challenging
	MedQA (41)	1,273	4	3 3
_	MedExpQA (6)	125	5	

Benchmarks (multiple choice question-answering)

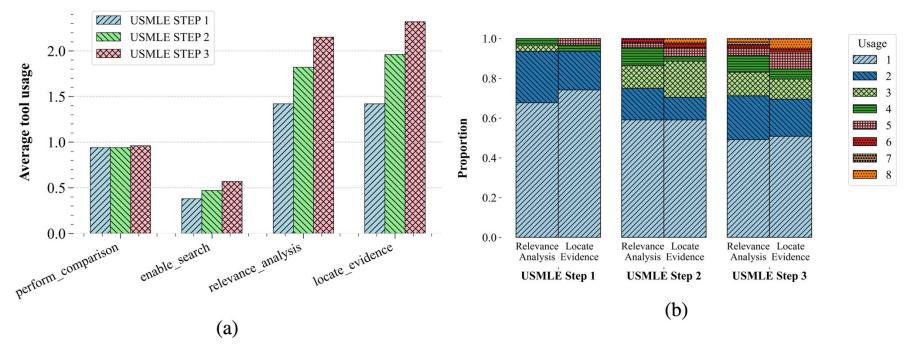
Model	USMLE Step 1	USMLE Step 2	USMLE Step 3	MedQA	MedExpQA
GPT-4	80.67	81.67	89.78	78.87	N/A
ChatGPT	51.26	60.83	58.39	50.82	N/A
BioMistral (7B)	34.04	37.61	37.70	41.01	37.60
OpenBioLLM (8B)	47.87	44.04	50.00	47.84	43.20
UltraMedical (8B)	42.55	27.52	34.43	38.49	35.20
OpenBioLLM (70B)	69.15	70.64	68.85	69.13	71.20
UltraMedical (70B)	70.21	55.05	56.56	52.32	50.40
PodGPT (70B)	73.40	72.48	74.59	65.04	63.20
Ours	82.98	86.24	<u>88.52</u>	<u>73.29</u>	78.40

Benchmarks (open-ended question-answering)

Benchmark	Model	BioMistral (7B)	OpenBioLLM (8B)	UltraMedical (8B)	OpenBioLLM (70B)	UltraMedical (70B)	PodGPT (70B)	Ours
	SFR	$0.79 \pm ext{0.09}$	0.70 ± 0.12	$0.81 \pm ext{0.13}$	0.85 ± 0.10	$0.82 \pm ext{0.11}$	0.86 ± 0.11	0.87 ± 0.09
USMLE Step 1	GTE	$0.48 \pm$ 0.17	$0.38 \pm$ 0.17	$0.57 \pm$ 0.21	$0.60 \pm$ 0.23	$0.63 \pm$ 0.23	$\overline{0.66}$ \pm 0.24	0.66 ± 0.22
-	BERTScore	$0.58 \pm$ 0.12	$0.51 \pm$ 0.13	0.61 ± 0.16	$0.66 \pm$ 0.17	$0.64 \pm$ 0.17	$\overline{0.68}$ \pm 0.20	0.68 ± 0.17
	SFR	$0.76 \pm ext{0.11}$	$0.71 \pm$ 0.10	0.80 ± 0.11	$0.82 \pm$ 0.09	0.80 ± 0.10	$\overline{0.85} \pm 0.10$	0.85 ± 0.09
USMLE Step 2	GTE	$0.45 \pm$ 0.19	$0.38 \pm$ 0.15	$0.52 \pm$ 0.19	$0.54 \pm$ 0.19	$0.59 \pm$ 0.22	$\overline{0.62}$ \pm 0.21	0.62 ± 0.22
	BERTScore	0.58 ± 0.11	$0.56 \pm$ 0.11	$0.61 \pm$ 0.13	$0.64 \pm$ 0.13	$0.63 \pm$ 0.14	$\overline{0.66} \pm 0.15$	0.67 ± 0.15
	SFR	$0.74 \pm ext{0.10}$	$0.70 \pm$ 0.10	$0.79 \pm ext{0.12}$	0.85 ± 0.11	$0.80 \pm$ 0.11	$\overline{0.84}$ \pm 0.11	0.86 ± 0.09
USMLE Step 3	GTE	$0.41 \pm$ 0.18	$0.38 \pm$ 0.14	$0.53 \pm$ 0.22	$\overline{0.63} \pm 0.26$	$0.60 \pm$ 0.23	$0.63 \pm$ 0.24	0.65 ± 0.22
	BERTScore	$0.57 \pm ext{0.11}$	$0.52 \pm$ 0.14	0.60 ± 0.17	$\overline{ extbf{0.71}} \pm ext{0.19}$	$0.62 \pm$ 0.15	$0.67 \pm$ 0.18	0.70 ± 0.17
	SFR	$0.76 \pm ext{0.10}$	$0.71 \pm$ 0.12	0.80 ± 0.12	0.86 ± 0.11	$0.80 \pm$ 0.11	$0.84 \pm$ 0.11	$\overline{0.85} \pm 0.10$
MedQA	GTE	$0.43 \pm$ 0.18	$0.40 \pm$ 0.17	$0.53 \pm$ 0.22	0.63 ± 0.26	$0.58 \pm$ 0.23	$0.60 \pm$ 0.23	$\overline{0.61}$ \pm 0.23
	BERTScore	$0.56 \pm$ 0.12	$0.52 \pm$ 0.15	$0.60 \pm$ 0.16	0.70 ± 0.19	$0.61 \pm$ 0.16	$0.65 \pm$ 0.18	$\overline{0.67}\pm$ 0.18
	SFR	$0.76 \pm ext{0.10}$	$0.71 \pm$ 0.11	$0.78 \pm$ 0.13	$0.81 \pm ext{0.11}$	$0.77 \pm$ 0.13	0.83 ± 0.11	$\overline{ extbf{0.84}} \pm ext{0.10}$
MedExpQA	GTE	$0.47 \pm$ 0.18	$0.40 \pm$ 0.18	$0.52 \pm$ 0.22	$0.54 \pm$ 0.24	$0.55 \pm$ 0.22	$\overline{\textbf{0.61}} \pm 0.23$	0.60 ± 0.22
	BERTScore	0.58 ± 0.11	$0.53 \pm$ 0.12	0.58 ± 0.15	$0.62 \pm$ 0.17	0.60 ± 0.14	0.65 ± 0.17	$\overline{\textbf{0.65}} \pm 0.16$

Credit: Jia et al., Agentic Memory-augmented Retrieval and Evidence Grounding in Medicine, Under Review.

Tool usage

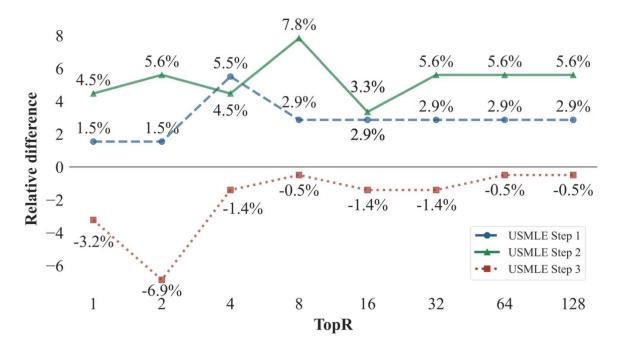


Credit: Jia et al., Agentic Memory-augmented Retrieval and Evidence Grounding in Medicine, Under Review.

Ablation study

Benchmark	USMLE Step 1	USMLE Step 2	USMLE Step 3	Average
Ours	82.98	86.24	88.52	85.91
w/o Tools	-1.07	-3.67	-4.91	-3.22
w/o Cache & Prune	-1.07	-2.75	-3.27	-2.36
w/o Evidence Search	-2.13	-3.67	-6.55	-4.12

Ablation study



Credit: Jia et al., Agentic Memory-augmented Retrieval and Evidence Grounding in Medicine, Under Review.

Ablation study

Table 1: Performance evaluation of different LLMs. We report different LLMs' performance (accuracy) across the USMLE Steps 1 to 3.

Model	USMLE Step 1	USMLE Step 2	USMLE Step 3
GPT-4	80.67	81.67	89.78
ChatGPT	51.26	60.83	58.39
Ours (Qwen2.5 72B)	<u>82.98</u>	86.24	88.52
Ours (Qwen3 32B)	91.49	87.16	86.07

Reasoning-based model

Runtime performance metric

Table 1: Latency (in seconds), per-GPU seconds per query (in seconds), and GPU hours per query (in hours) of the agentic system deployed using 4 NVIDIA L40S GPUs.

Qwen2.5 72B	USMLE Step 1	USMLE Step 2	USMLE Step 3
Latency	1.12	1.48	1.80
per-GPU seconds per query	39.15	51.64	83.30
GPU hours per query	0.043	0.057	0.093

Take away

- Agentic framework unifies document retrieval, evidence grounding, as well as Al generation, with an automatic and dynamic process
- Tool-augmented LLM-based agent enables dynamic multistep tool use, eliminating the need for manually engineered prompts or multi-stage pipelines
- The cache-and-prune memory bank mechanism efficiently extends the retention of relevant documents for evidence grounding, enhancing diagnostic accuracy and computational efficiency

Thank you very much for your attention!

