

Real World Applications of Generative AI in Liver Diseases and Transplantation

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CPH 100 Guest Lecture

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Disclosures

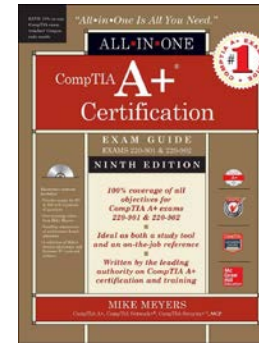
- **Research funding:**
 - NIH NIDDK K23DK139455
 - AGA Research Pilot Award
 - Gilead Sciences Investigator Initiated Funding
- **I disclose the following financial relationship(s) with a commercial interest within the last 48 months:**
 - Merck and Co (Grant, Concluded)
 - Madrigal Pharmaceuticals (Consultant)
 - Gilead Sciences (Advisory Board, Consultant)

- **I DID use artificial intelligence to prepare portions of this presentation, and I take full responsibility for the contents in this presentation**

About Me

- Assistant Professor of Medicine, In Residence
- Gastroenterologist/Transplant Hepatologist
- Divisions of Gastroenterology/Hepatology and Clinical Informatics and Digital Transformation
- Joined UCSF Faculty in July 2022
- AGG Faculty in CPH
- Affiliate Faculty in Bakar Computational Health Sciences Institute
- NIDDK K23 Career Development Award

**2001:
CompTIA A+ Certification**



**2008-2010:
Boston Consulting Group**



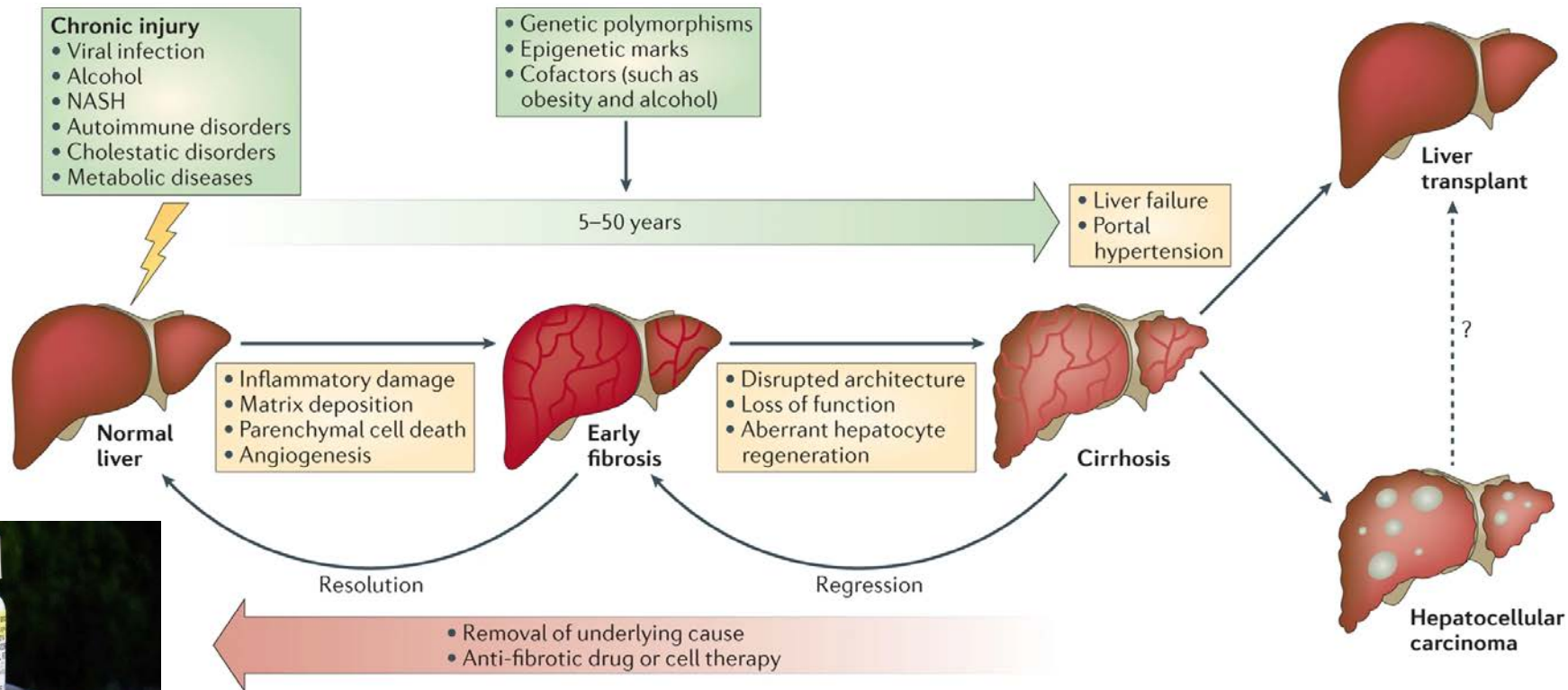
**2005:
Residential Computing Consultant
at Princeton**



**2014:
Digital Health Startup**



Liver Disease is Complicated



Nature Reviews | Immunology

Why AI in Hepatology and Liver Diseases?

Liver Disease Burden

200,000+

Hospitalizations for cirrhosis annually

\$20 billion

In annual spending

37%

Readmitted within 30 days

The GenAI Opportunity

Unstructured data

- ~80% of clinical data is in free-text notes — LLMs can process this at scale

Complex decision-making

- Transplant selection, treatment choice, risk stratification

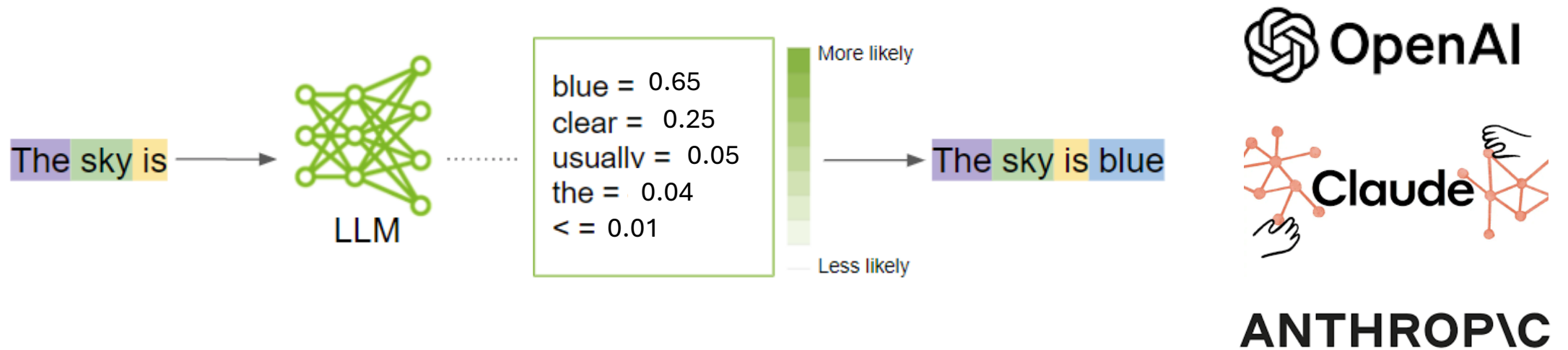
Guideline adherence gaps

- AI-powered decision support could close the evidence-to-practice gap

2022: ChatGPT

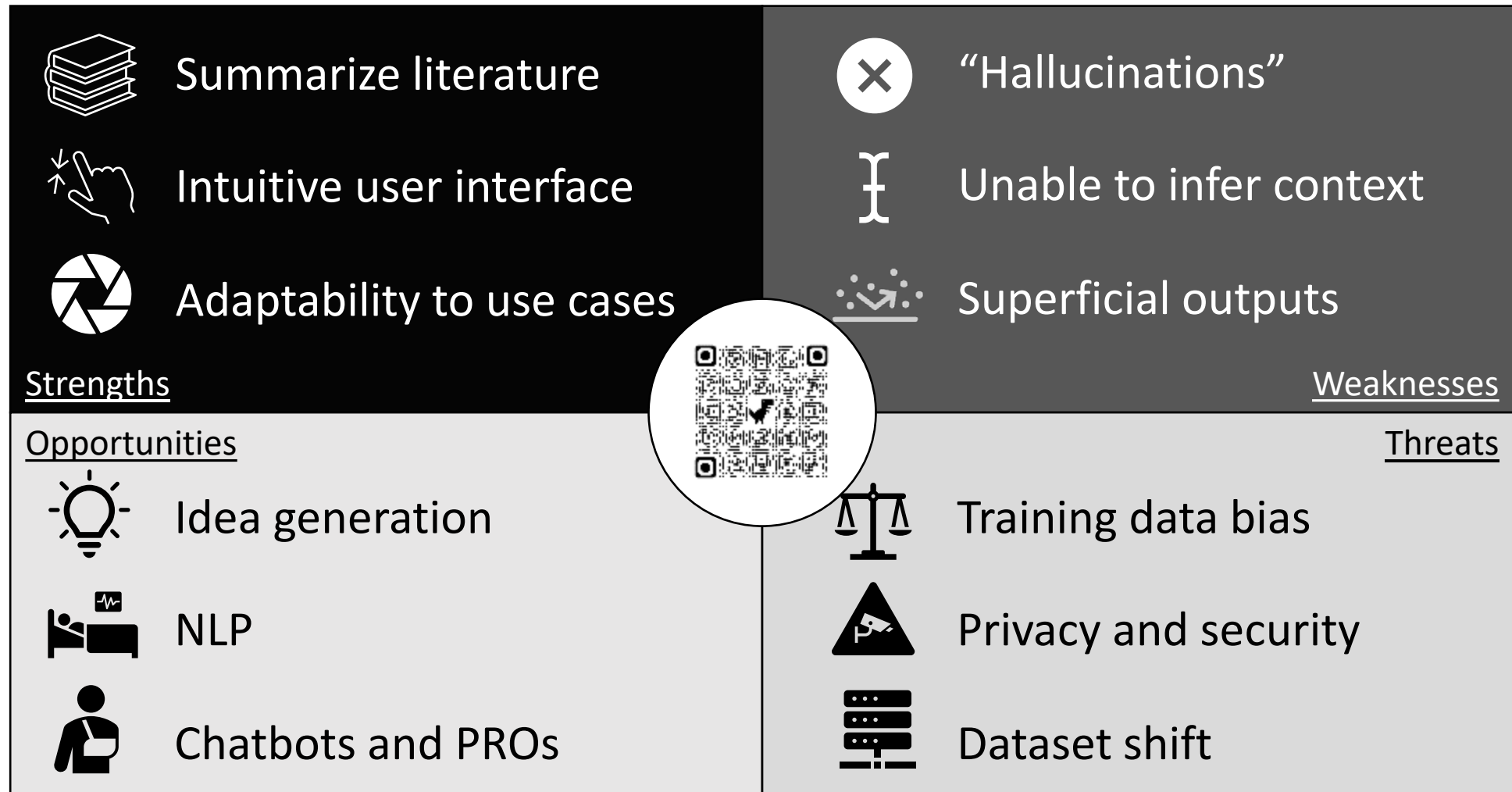


Large Language Models (LLMs) predict and generate human-like responses



- LLMs process text using neural architectures trained through statistical pattern recognition across massive datasets
- Like a super-powered autocomplete, LLMs predict what words should come next

2022: Potentials and pitfalls of ChatGPT



Motivating Examples



How can we use GenAI to derive new insights from the clinical record?



How can LLMs help us run, and learn from, a clinical trial?



How can LLMs assist with clinical decision making?

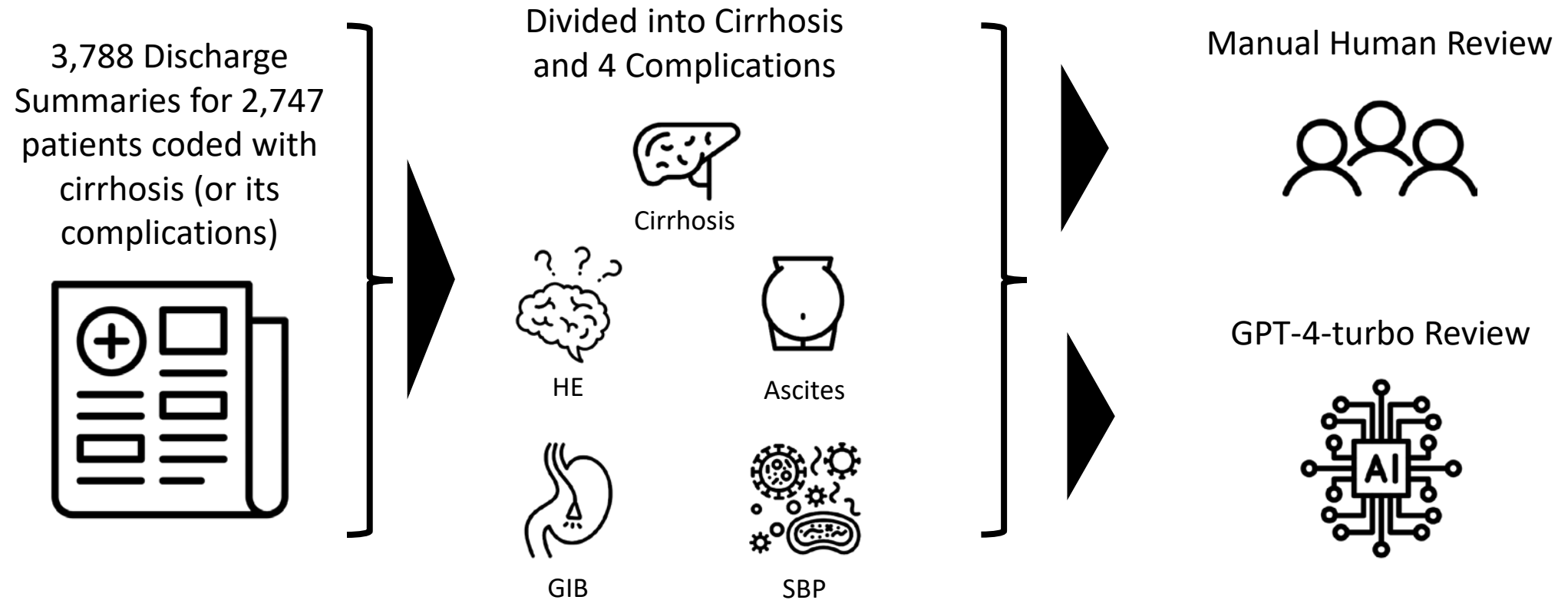


How do we use GenAI to construct novel diagnostic tests?

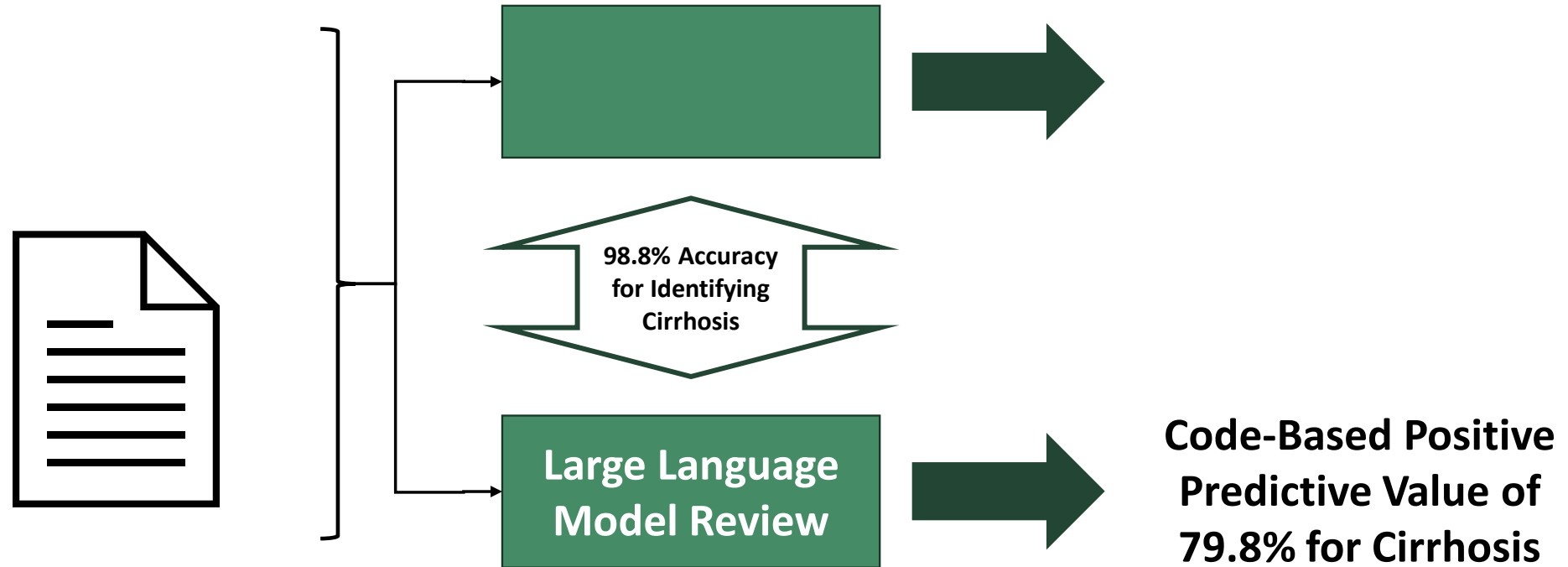


How can we use GenAI to derive new insights from the clinical record?

Cirrhosis cohort identification in discharge summaries



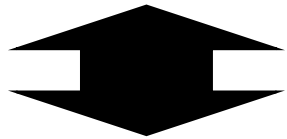
Code-based methods had PPVs of ~80% versus humans and LLMs



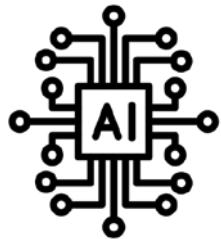
PPVs for complications of cirrhosis were much lower: 54% for HE, 55% for ascites, 68% for GIB, and 66% for SBP

Humans may not be better than LLMs?

Manual Human Review



GPT-4-turbo Review



	# Notes	LLM Accuracy	Human "Error" Rate
Cirrhosis	3,788	98.2-98.8%	4.3%
HE	1,969	94.4-96.3%	14.8%
Ascites	1,618	91.4-94.4%	18.5%
GIB	1,127	85.4-87.8%	11.0%
SBP	339	90.7%	11.6%

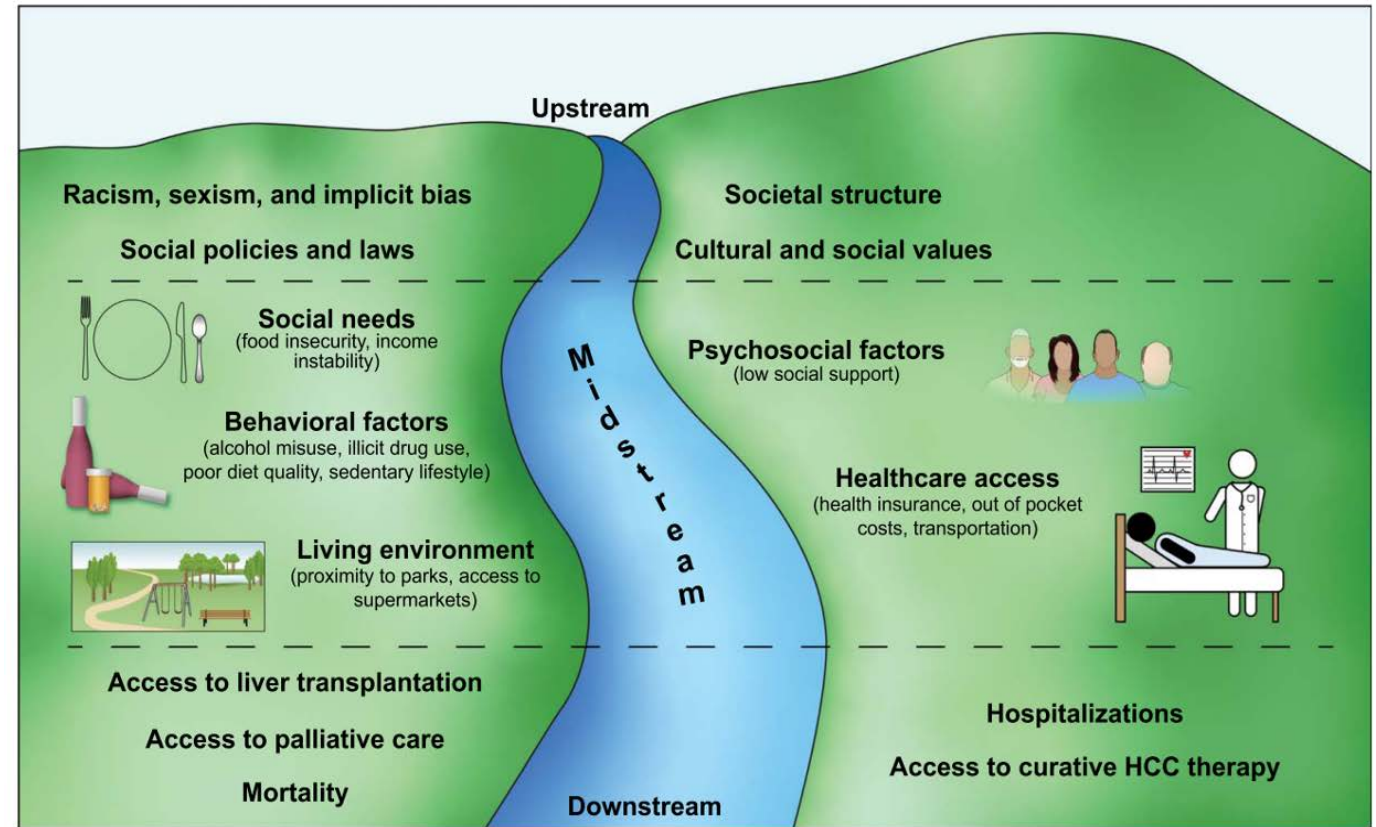
Psychosocial risk factors are associated with liver transplant outcomes

Impact on waitlist registration

- Psychosocial risk and SDOH factors disproportionately concentrate liver disease risk factors (e.g., alcohol use, opioid use, obesity) among historically disadvantaged populations

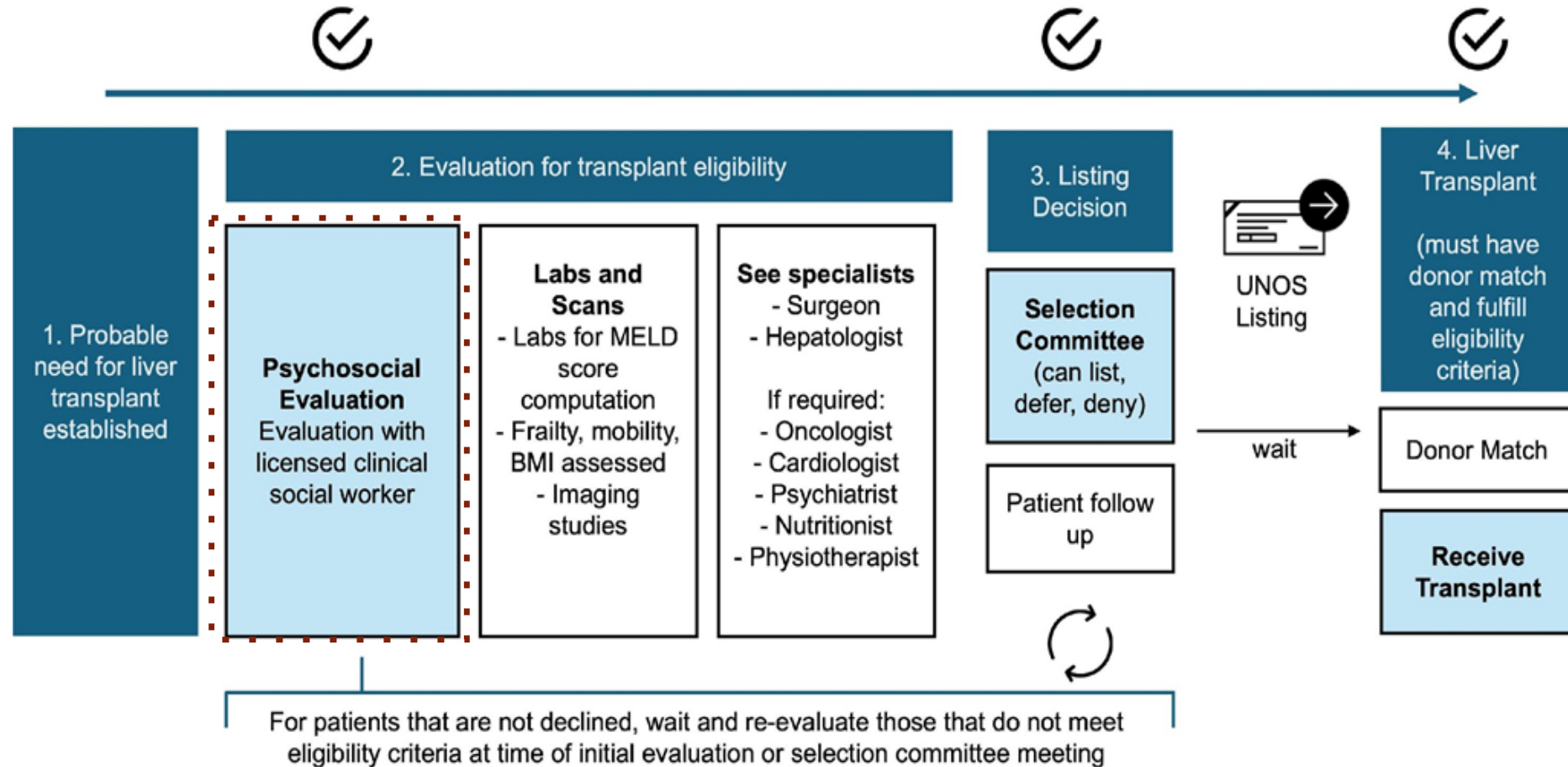
Impact on successful transplant

- Disparities persist through the transplant process, with women and racial and ethnic minorities facing reduced access to evaluation, listing, and transplantation

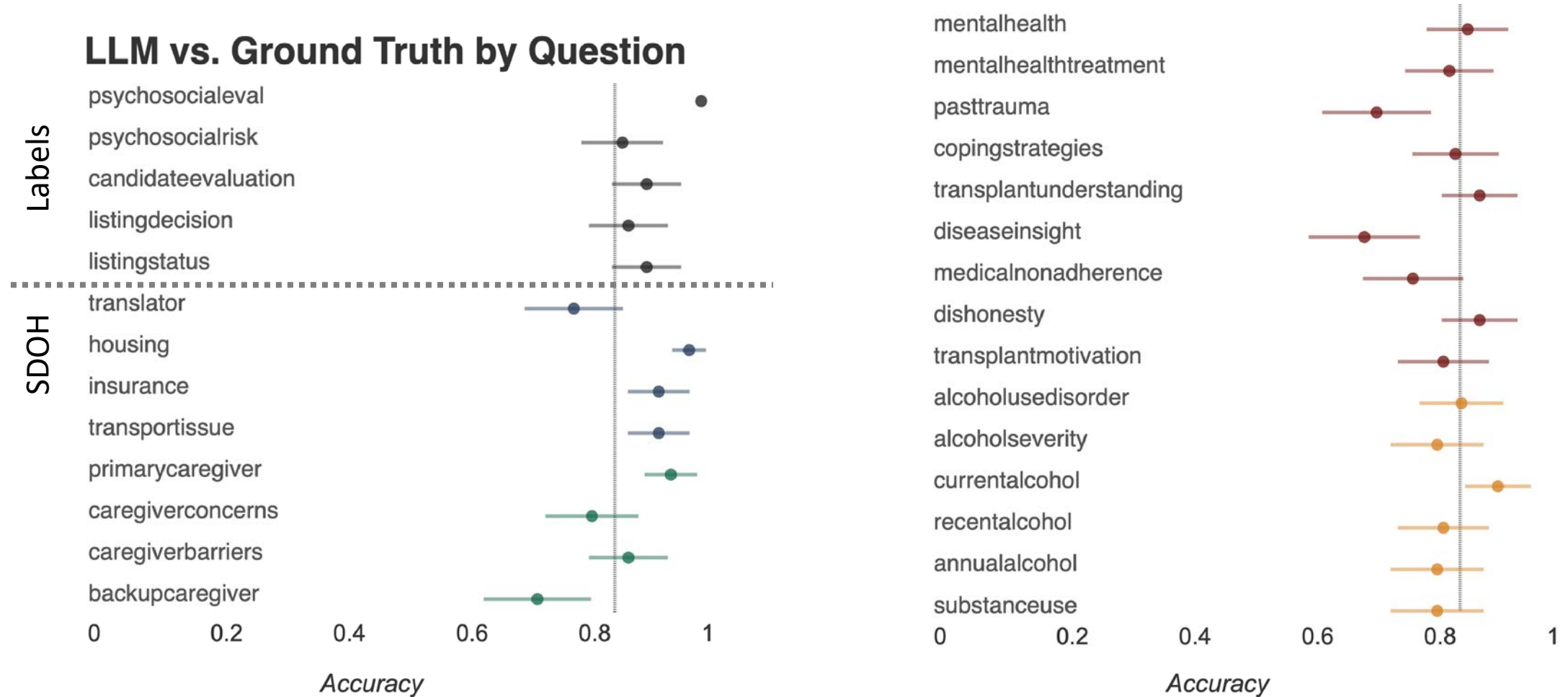


Contribution of Structural and psychosocial/SDOH factors to Liver Disease Outcomes (from Kardashian et al. 2022)

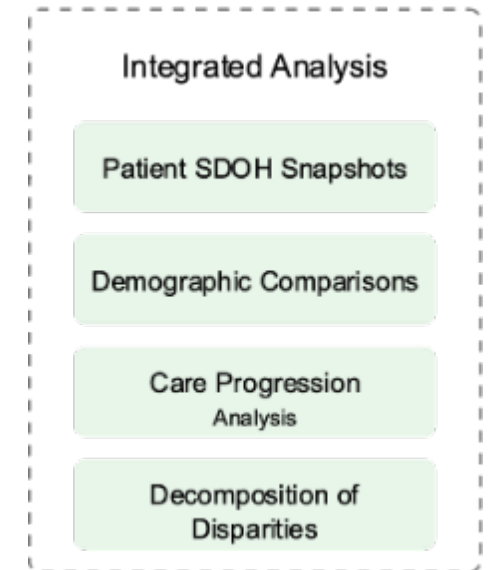
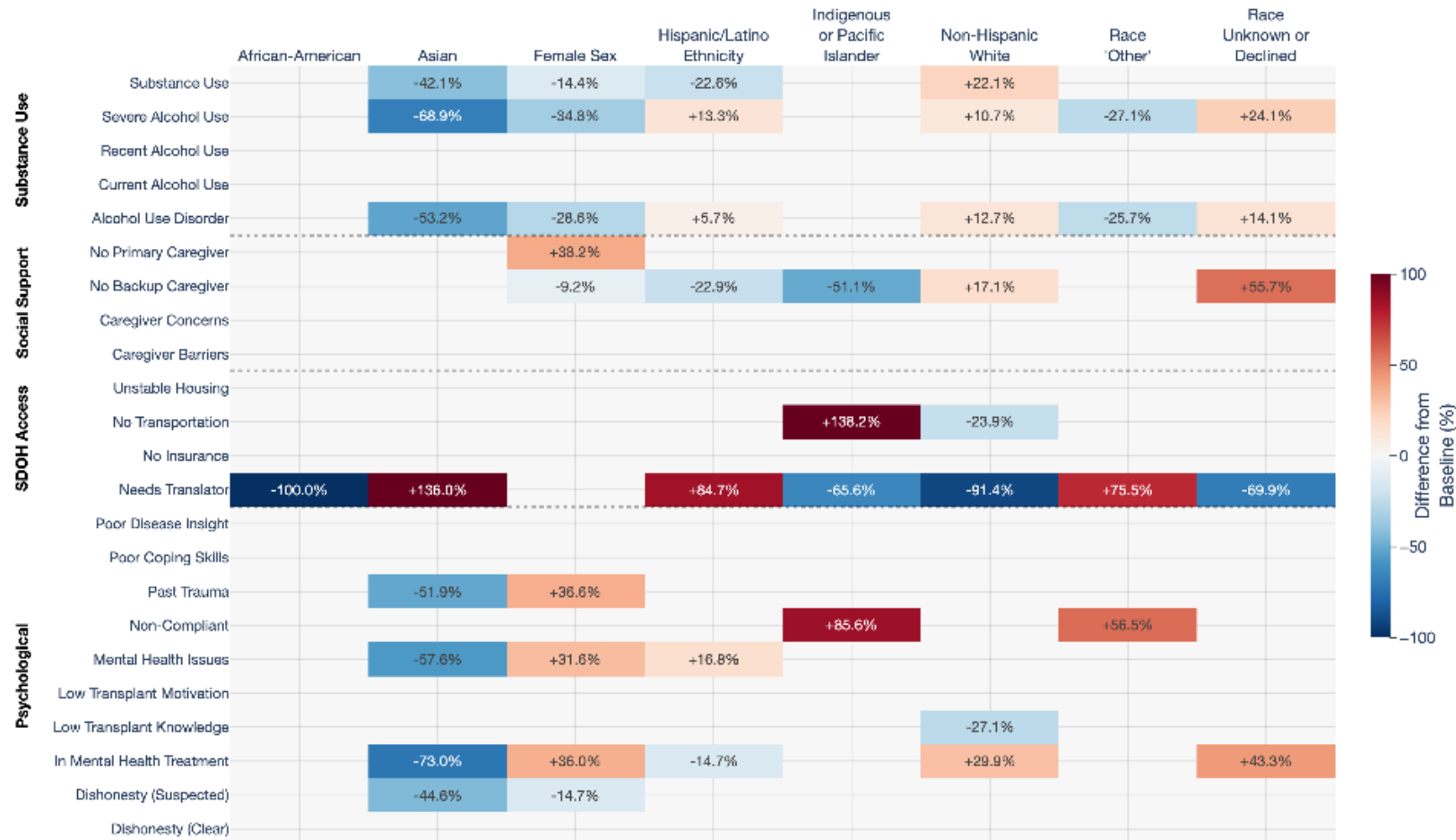
LT psychosocial evaluation notes systematically capture these factors



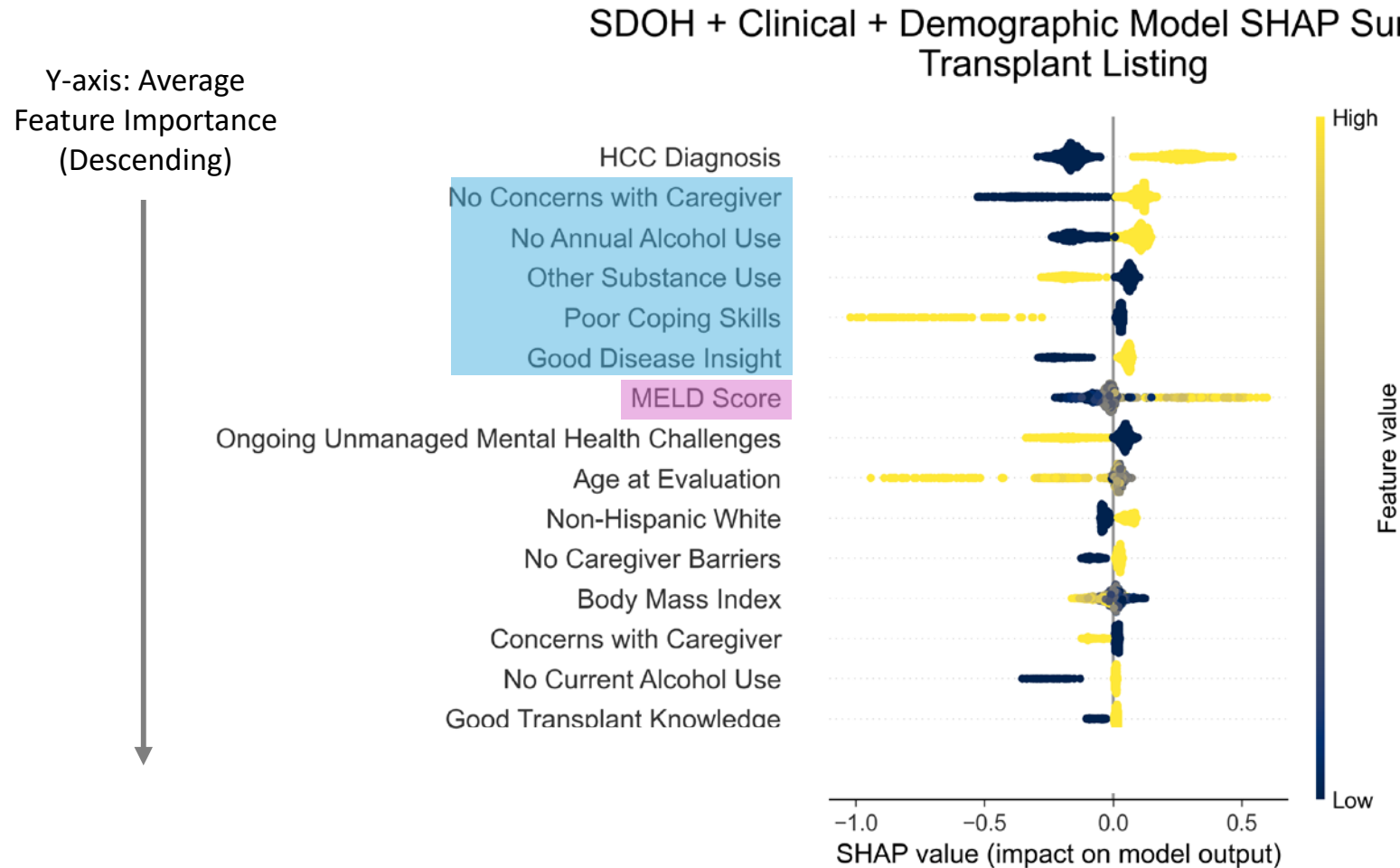
LLMs extracted psychosocial risk and SDOH factors with 86% accuracy



Psychosocial risk and SDOH categorization from SW evaluation notes



Psychosocial/SDOH factors may be *more important than* MELD for LT listing





How can LLMs help us run, and learn from, a clinical trial?

CirrhosisRx: A stepped-wedge cluster randomized trial of a CDS for cirrhosis care



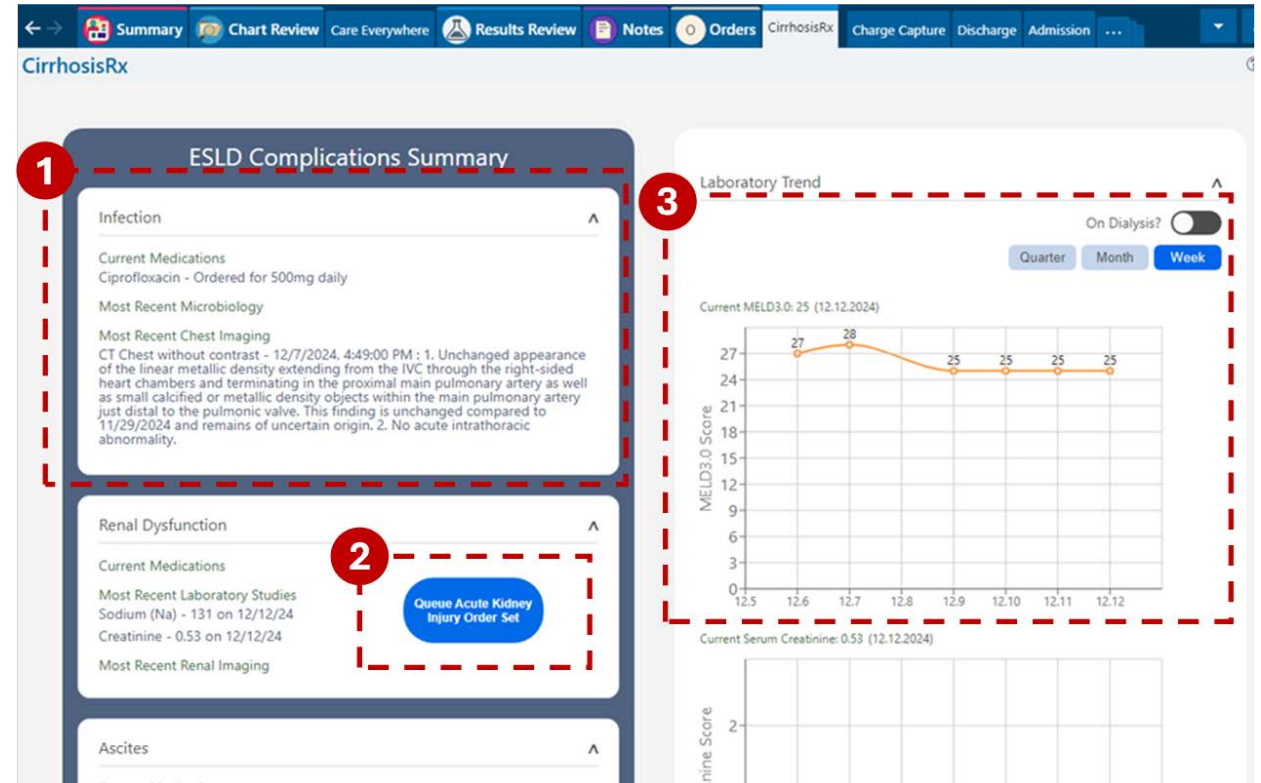
Step	Months 1-3	Months 3-6	Months 6-9	Months 9-12	Months 12-15	Months 15-18	Months 18-21	Months 21-24
1								
2								
3								
4								
5								
6								
7								
8								

CirrhosisRx Assignment

"Usual Care" Assignment

- **Intervention and primary outcome**

- SMART-on-FHIR CDS in Epic surfacing AGA/AASLD process measures for ascites, SBP, HE, and GI bleeding
- Primary: Opportunity-level adherence to 7 measures, contraindication-netted



Trial flow and analytical approach

- **Primary intention-to-treat model**
 - Mixed-effects logistic at the opportunity level: intervention + calendar period + MELD
 - Random intercepts: cluster, treatment team, patient
- **Robustness (pre-specified)**
 - Adjustment waterfall: crude => +period => +team => +MELD
 - Permutation (randomization) inference
 - Missed-opportunity count and proportion on the randomization-fixed denominator
 - Per-protocol scenarios and complier-average causal effect (IV)
 - Two code-sets (python and R for replication and cross-checking)

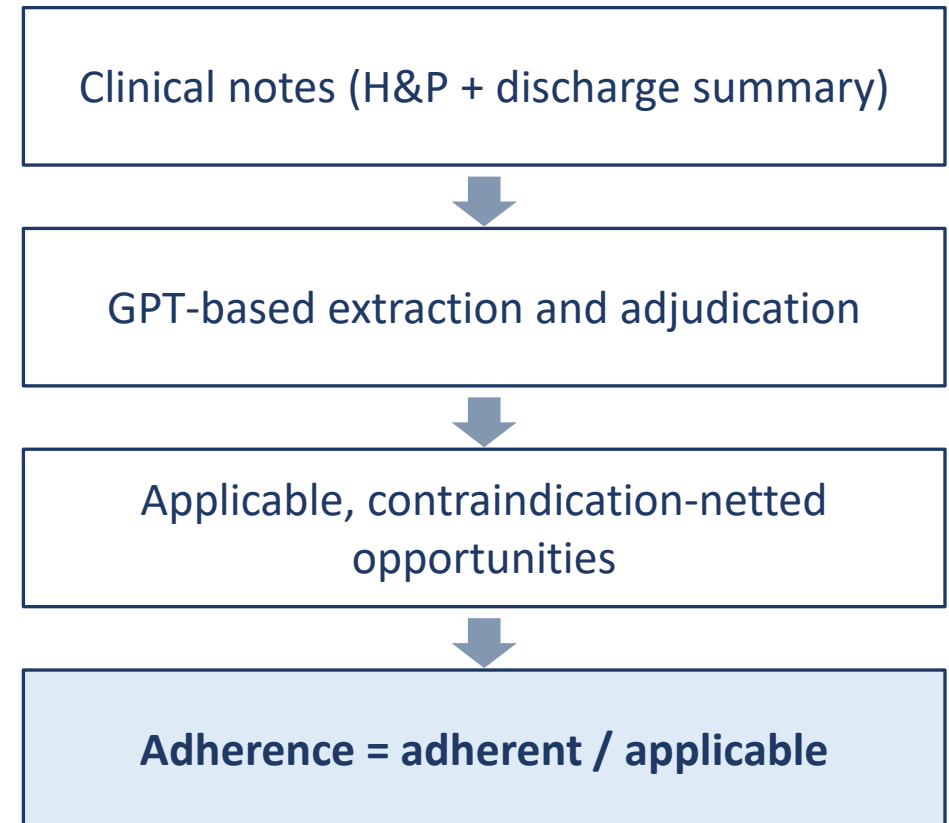
LLMs define the cohort, the applicable measures, and the contraindication-netted endpoint

- **Three ascertainment tasks done by AI**

- Cohort: H&P-suspected cirrhosis => analytical dataset (denominator)
- Guideline applicability: Which of the 5 guideline measures for inpatients with cirrhosis are applicable (complications treated during the admission ascertained from discharge summary)
- Netting: Opportunities with documented contraindications removed from the denominator (e.g., a patient without ascites who is indicated for a paracentesis)

- **Why use an LLM?**

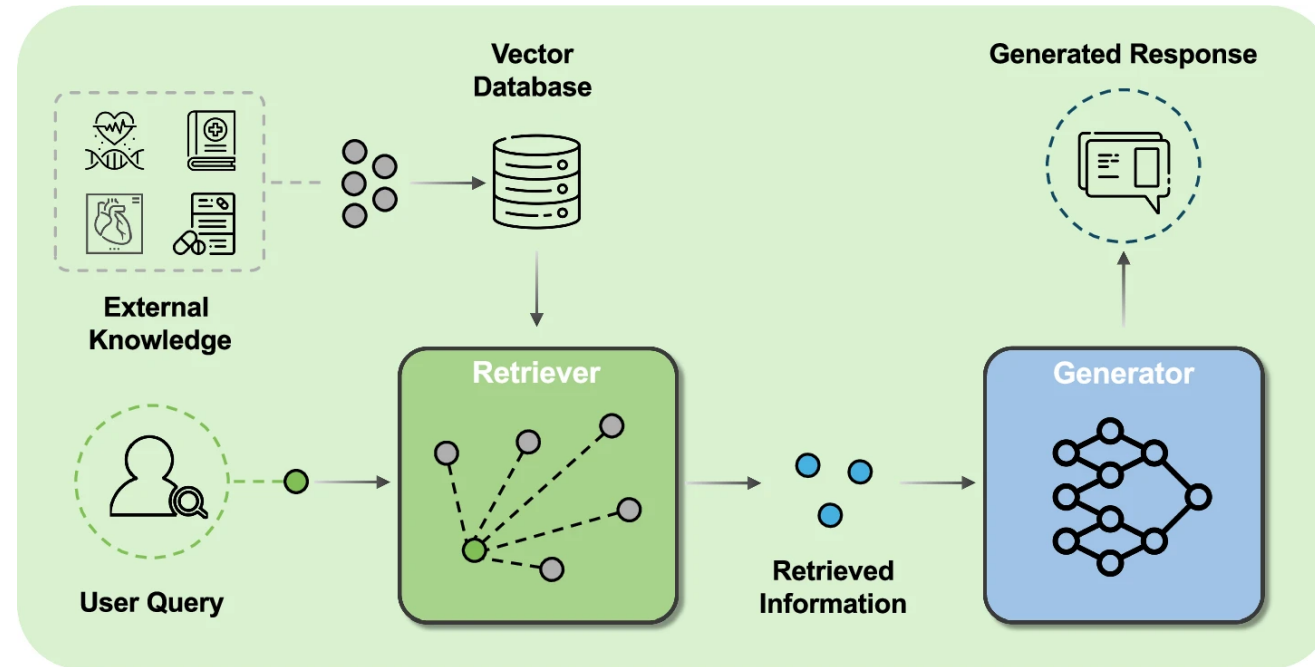
- ICD-10 codes are not accurate
- Tens of thousands of admissions × note-level adjudication (beyond manual review)





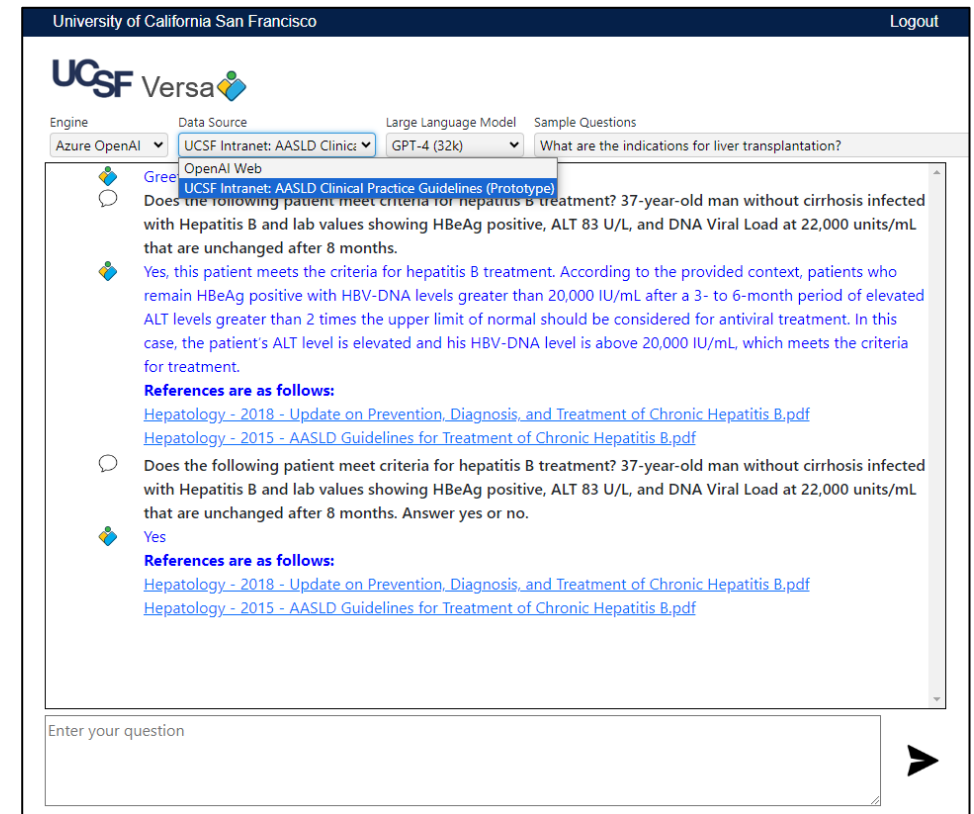
How can LLMs assist with clinical decision making?

Retrieval augmented generation allows for customized LLMs



- External data is first encoded into vectors and stored in the vector database (where vectors are mathematical representations of various types of data in a high-dimensional space).
- In the retrieval stage, when receiving a user query, the retriever searches for the most relevant information from the vector database. In the generation stage, both the user's query and the retrieved information are used to prompt the model to generate content.

LiVersa – A customized liver-disease specific large language model



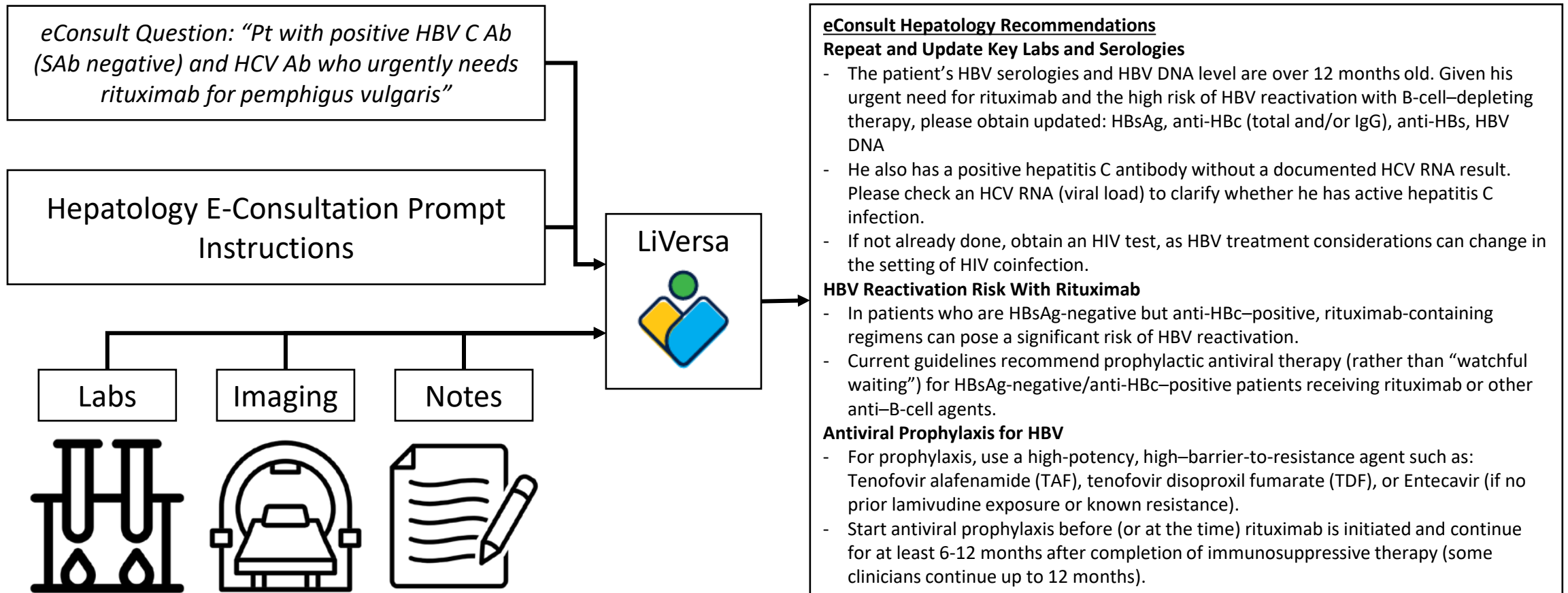
LiVersa matched ChatGPT and outperformed LLaMA in answering hepatology questions

Human-Evaluations of GenAI/LLM Answers to 10 Hepatology Topic Questions

				
	LiVersa vs. LLaMA		LiVersa vs. ChatGPT	
Accuracy	4	vs. 0	2	vs. 1
Comprehensiveness	5	vs. 4	0	vs. 3
Safety	3	vs. 1	2	vs. 3

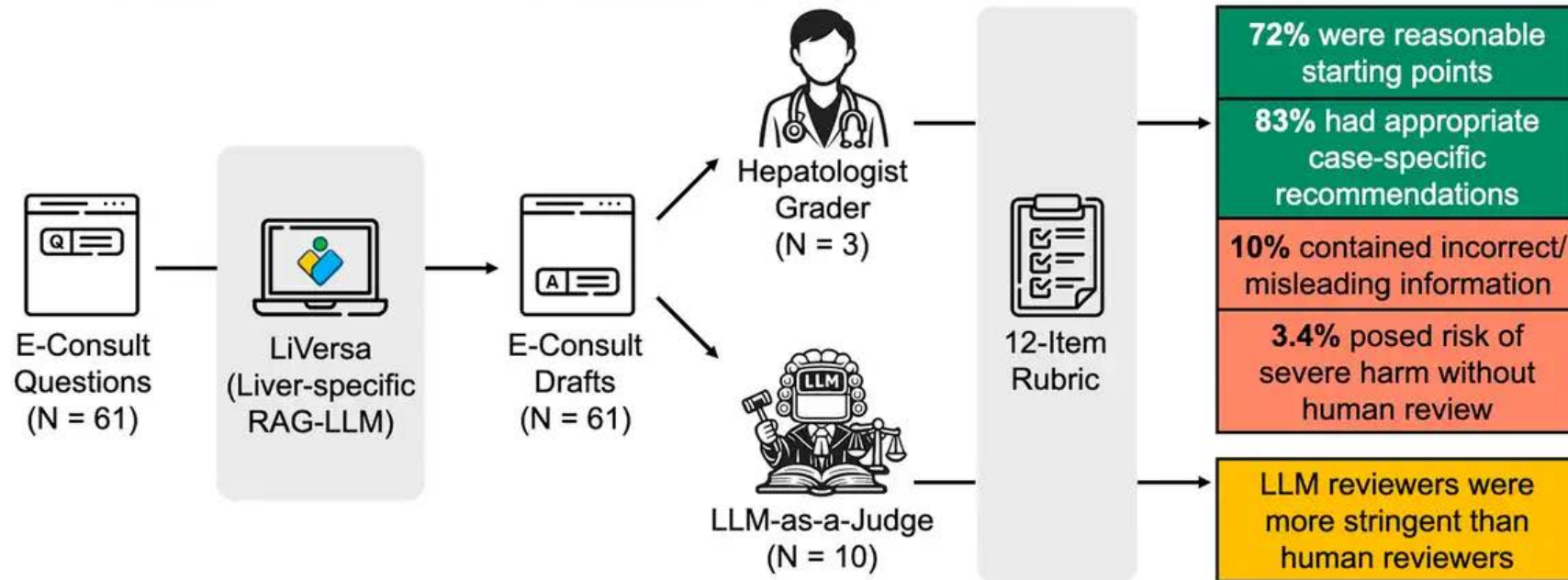
LLM	PSC Question - What is most common type of malignancy associated with Primary Sclerosing Cholangitis?	Character Count
LiVersa	The most common type of malignancy associated with Primary Sclerosing Cholangitis (PSC) is cholangiocarcinoma, which is a cancer that arises from the bile ducts.	161
ChatGPT 4	The most common type of malignancy associated with Primary Sclerosing Cholangitis (PSC) is cholangiocarcinoma, a cancer of the bile ducts. This cancer occurs more frequently in individuals with PSC than in the general population. The risk of developing cholangiocarcinoma in patients with PSC is significantly elevated, making regular monitoring and early detection vital. Additionally, PSC patients are at an increased risk for other cancers, including colorectal cancer, particularly in those with concurrent inflammatory bowel disease. Early diagnosis and management of PSC are crucial to monitor and potentially reduce the risk of these malignancies.	654
LLaMA 2	The most common type of malignancy associated with Primary Sclerosing Cholangitis (PSC) is cholangiocarcinoma, which is a rare cancer that originates in the epithelial cells lining the bile ducts. PSC increases the risk of developing cholangiocarcinoma by up to 30-fold compared to the general population. Other types of cancer that can occur in association with PSC include gallbladder cancer and pancreatic cancer. The exact mechanisms by which PSC leads to malignancy are not fully understood but may involve chronic inflammation and genetic mutations. Early detection and treatment of PSC can help reduce the risk of developing malignancies.	645

Specialty-specific LLMs integrated into EHR for consultation drafting (LiVersa)



Retrospective testing showed that LiVersa could generate accurate e-consult drafts

Hepatology e-consult responses generated by artificial intelligence demonstrate accuracy but require human oversight



We took a broad cross-section of actual e-consults completed over 3 months

TABLE 1 Characteristics of e-consult requests (n = 61)

Characteristic	
Patient demographics	
Sex, n (%)	
Female	43 (70.5)
Male	18 (29.5)
Age, years	
Mean	55.0
Median (range)	54.0 (23–97)
E-consult category, n (%)	
Abnormal liver function tests	21 (34.4)
Hepatitis B	14 (23.0)
Abnormal imaging	13 (21.3)
FibroScan	6 (9.8)
Hepatitis C	4 (6.6)
Steatosis	3 (4.9)
Clinical question characteristics	
Word count, mean (SD)	77 (65)
Character count, mean (SD)	484 (420)
Word count by category, mean (SD)	
Abnormal imaging	109 (51)
Abnormal liver function test	97 (86)
Steatosis	75 (57)
Hepatitis C	48 (39)
Hepatitis B	45 (26)
FibroScan	29 (21)

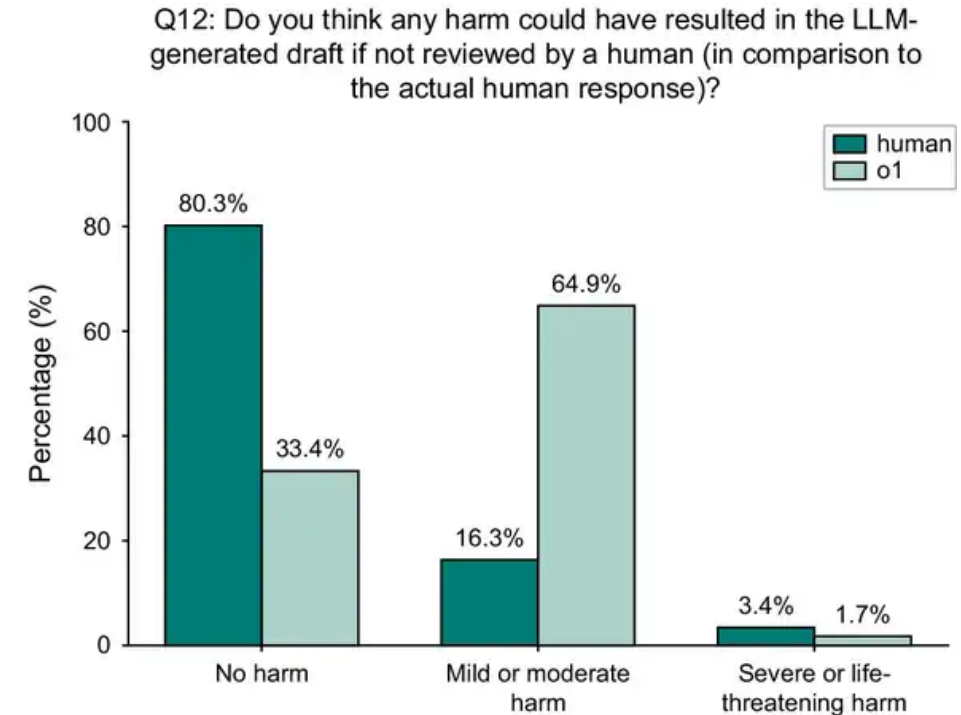
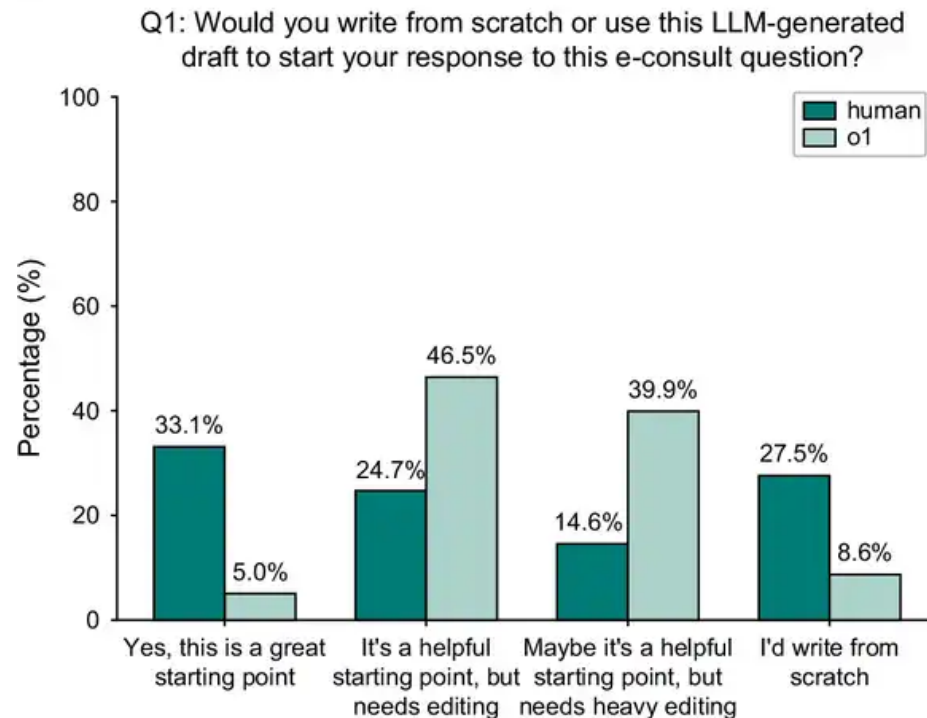
Abbreviation: e-consult, electronic consultation.

TABLE 2 Characteristics of e-consult responses by hepatologists versus LiVersa (n = 61 pairs)

Characteristic	Hepatologist	LiVersa	p
Overall response characteristics, mean (SD)			
Word count	284 (172)	264 (146)	0.47
Character count	1765 (1136)	1798 (1038)	0.86
Sentence count	14 (14)	11 (7)	0.10
Words per sentence	24 (13)	25 (6)	0.44
Word count by category, mean (SD)			
Abnormal imaging	367 (298)	282 (186)	0.35
Abnormal liver function tests	277 (115)	286 (158)	0.84
FibroScan	204 (60)	218 (99)	0.73
Hepatitis B	258 (94)	240 (113)	0.67
Hepatitis C	160 (125)	283 (103)	0.01
Steatosis	413 (82)	217 (204)	0.23
Words per sentence by category, mean (SD)			
Abnormal imaging	23 (17)	28 (4)	0.37
Abnormal liver function tests	26 (11)	25 (6)	0.84
FibroScan	16 (5)	17 (3)	0.58
Hepatitis B	24 (9)	27 (4)	0.22
Hepatitis C	17 (1)	27 (3)	0.01
Steatosis	42 (19)	21 (4)	0.24

Note: p-values are from paired t tests (by case) comparing hepatologist versus LiVersa responses to each e-consult question.
Abbreviation: e-consult, electronic consultation.

72.5% of AI generated e-consult drafts were “good enough” as starting points

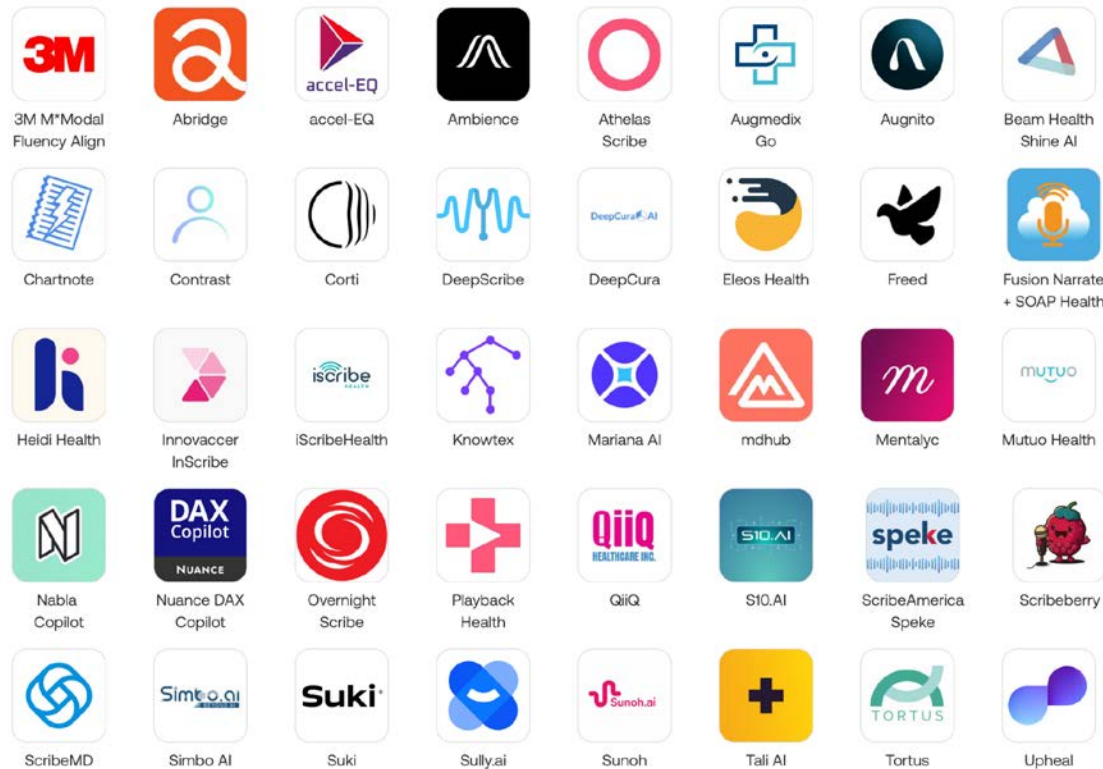


LiVersa E-consults await active integration into the UCSF EHR for a prospective randomized controlled trial



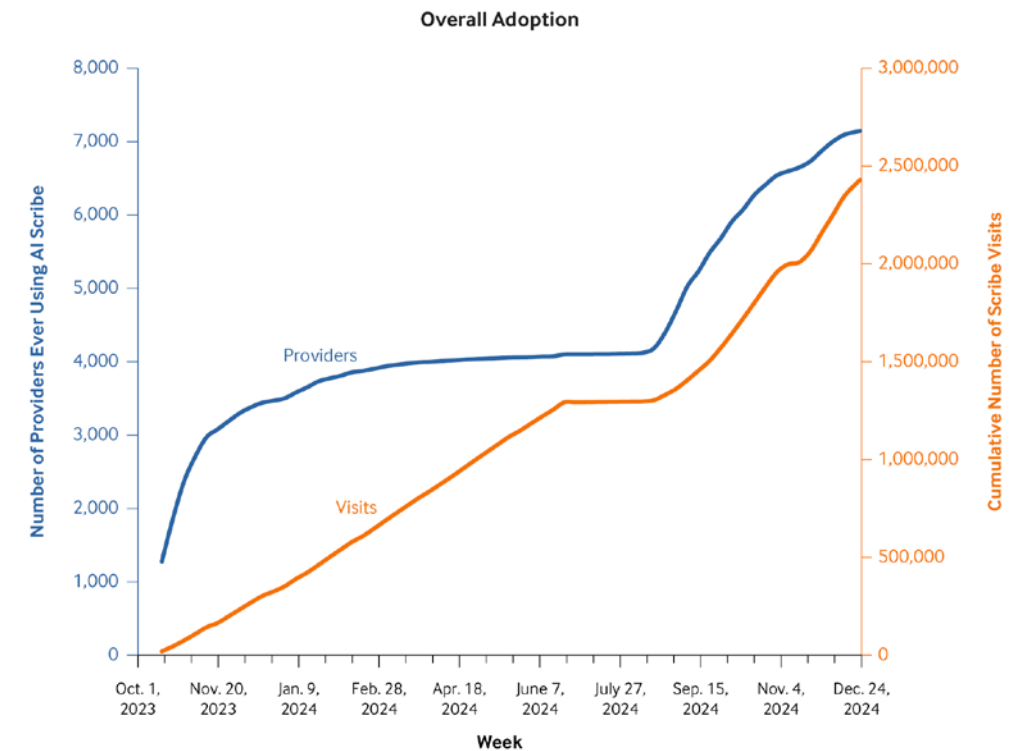
How do we use GenAI to construct novel diagnostic tests?

Ambient AI scribes are rapidly being adopted



Growth in AI Scribe Usage

This chart shows the overall adoption of providers using an AI scribe and the cumulative number of AI scribe visits, from October 2023 through December 2024.



Source: The authors
NEJM Catalyst (catalyst.nejm.org) © Massachusetts Medical Society

<https://elion.health/resources/ai-ambient-scribes>
Tierney et al. NEJM Catalyst, 2025.

Covert hepatic encephalopathy (CHE) is common, costly, and clinically consequential

40.9%

Pooled Prevalence

Up to 50% of all patients
with cirrhosis affected

25% Child-Pugh A →
52% Child-Pugh C

Unmet Need



Reversible

With Lactulose + rRifaximin

50%

Ever Screened

Clinical Consequences



5.8×

Risk of motor vehicle
accidents



HR 2.5

Risk of hospitalization



HR 3.4

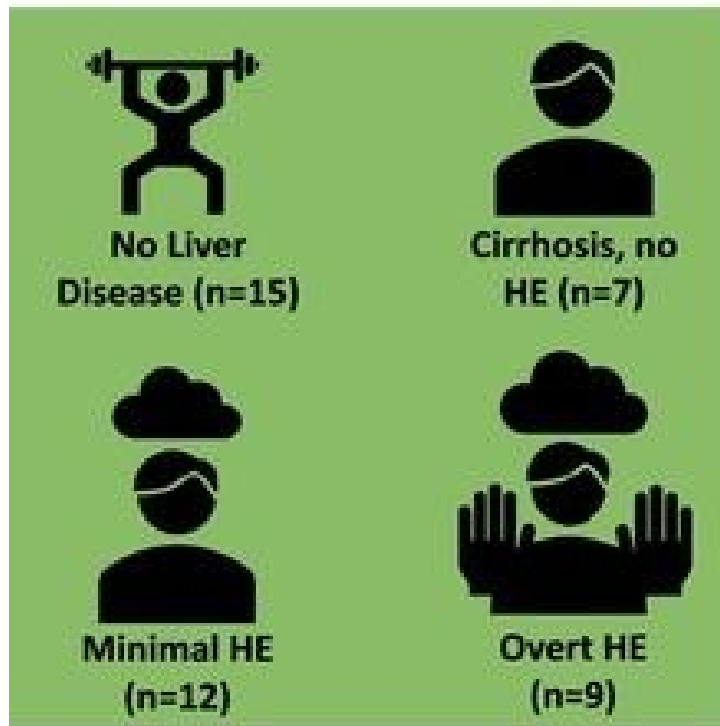
Risk of mortality



\$11.9B

Annual US HE-related
charges

SPEED: Speech patterns and enunciation for encephalopathy determination



Recordings taken of participants reading sentences, a standardized passage, and completing the Cookie Theft Picture



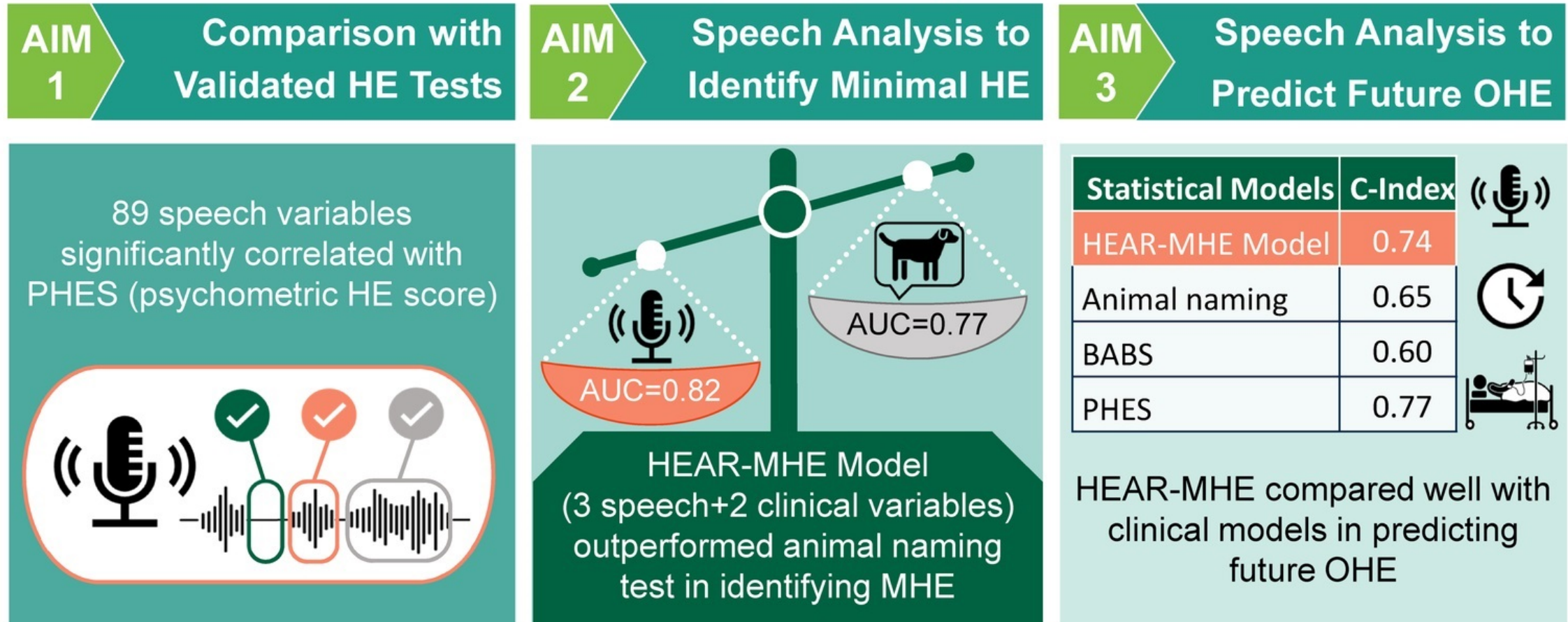
Speech rate, precision, and overall severity measured by blinded trained analysts



- (1) Speech rate, precision, and overall severity were worse in overt HE compared to minimal HE and no HE
- (2) As overt HE improved, speech rate and precision improved
- (3) Minimal HE patients had impairments in speech precision and overall severity compared to no HE

Speech could be used to monitor HE severity remotely or in hospitalized patients

HEAR-MHE: Automated speech analysis for covert hepatic encephalopathy



Speech carries the CHE signal, but scripted elicitation captures only part of it



Standard Reading Passages

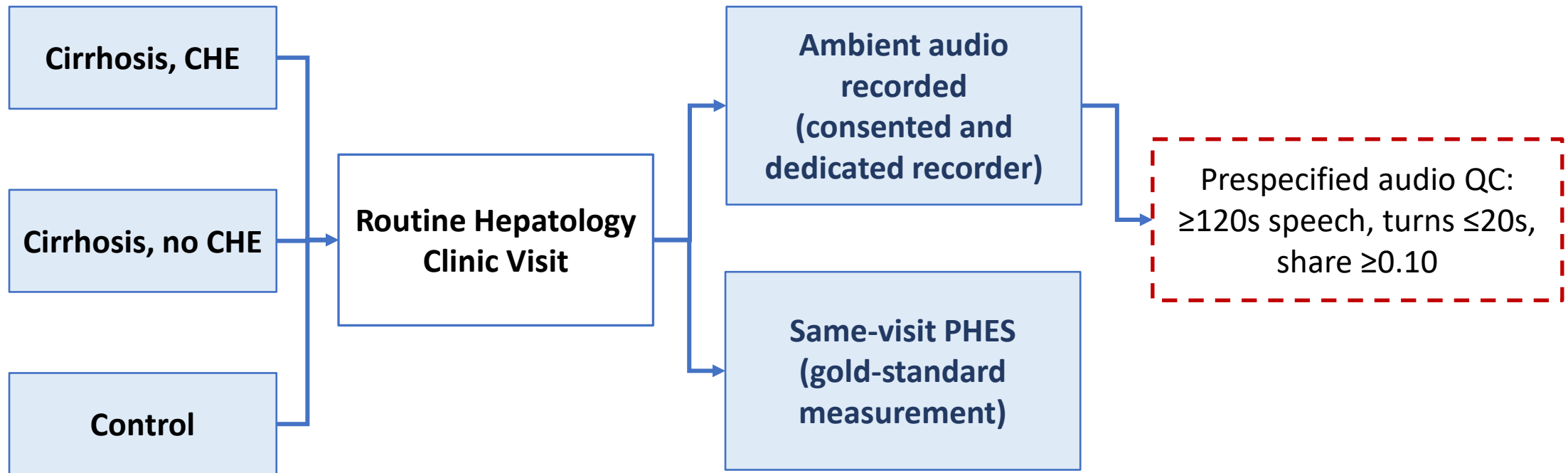
- Acoustic signals only
- Requires literacy
- Dedicated testing time
- Cross-sectional snapshot



Spontaneous Clinical Conversations

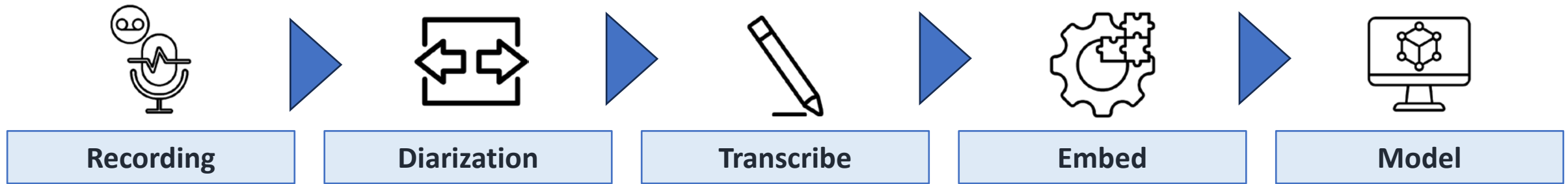
- Acoustic + lexical/semantic signals
- No literacy or language prerequisites
- Captured passively during routine care
- Longitudinal trajectory over time

Three-arm pilot study to associate ambient audio recordings to gold standard measures



Reference standard and specimen come from the same visit — no added patient burden

Analytical pipeline relies upon LLM conversion of audio to transcript to embeddings



Timing / Psychomotor

Response latency, turn duration, talk share
(conversational analogue of psychomotor testing)

Discourse Coherence

Adjacent-turn similarity, question–answer relevance,
coherence baseline (measurable only in spontaneous
dialogue)

Lexical

Vocabulary diversity (MTLD), content-to-function ratio,
sentence length, dependency depth

Acoustic (Voice)

Pilot: pyAudioAnalysis, 136 surfaced & tiered
(chroma/deltas)

Research Roadmap for Ambient-CHE



Pilot Study (Current)

20-30 participants
Feasibility + effect size
Proof-of-concept



Confirmation Study (Next)

600 participants
Longitudinal data



EHR-Integrated CDS

SMART-on-FHIR app
Silent study → pRCT



Clinical Impact

Passive CHE detection
Therapy initiation
OHE prevention

Future state: Ambient AI scribes will not only transcribe encounters but also provide validated diagnostic information, seamlessly and without workflow disruption

Conclusions



- **Reading the record:** LLMs read thousands of notes as accurately as humans making manual chart review functionally obsolete



- **Accelerating research:** LLMs can define the trial cohort and endpoints, draft analytical code, and work in tandem with humans to catch bugs and interpret results



- **Drafting, not deciding:** LiVersa, a liver-specific, privacy-protected LLM, wrote e-consult drafts that were “good enough” starting points in 72.5% of the time



- **New diagnostics:** Routine clinic conversation captured by ambient recording carries a measurable signal for covert hepatic encephalopathy



- **For the next generation of clinicians and scientists:** the open problems are data quality, workflow, trust, and rigorous evaluation

A Tip for the Future



Questions?

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