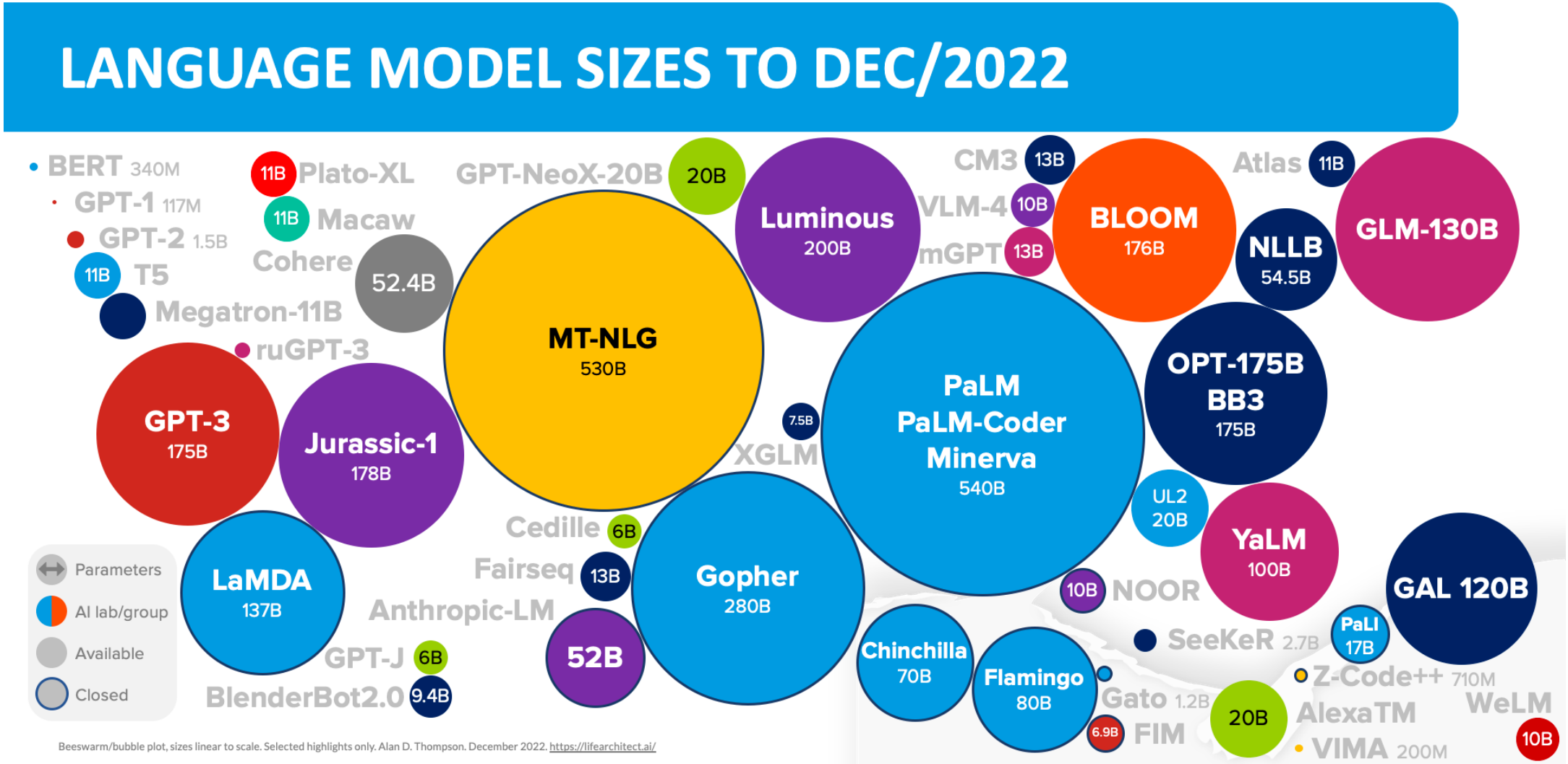
上海人工智能实验室
Shanghai Artificial Intelligence Laboratory

Foundation Models in Medicine Generalist vs Specialist

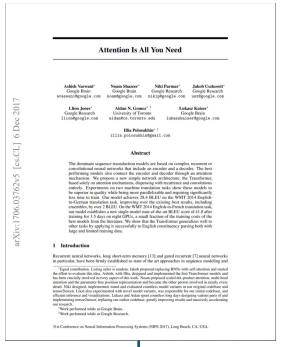
Presented by Shaoting Zhang

Date：2023.12

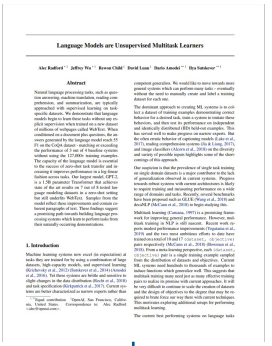
Large-Language Models (LLM)



Large-Language Models (LLM)


Attention Is All You Need


BERT: Pre-training of Deep Bidirectional Transformers for Language Understanding


Language Models are Unsupervised Multitask Learners


GPT-4 Technical Report

OpenAI

2017/06 Transformer

- Proposed the Transformer architecture

2018/06 GPT

- Using the decoder of Transformer, a language model is trained on a large amount of unannotated text data, which is then fine-tuned for subtasks

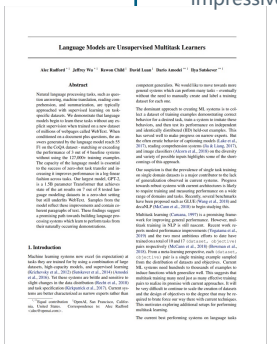

Improving Language Understanding by Generative Pre-Training

2018/10 BERT

- Using the encoder of Transformer and pre-training on a larger dataset, the performance is much better than GPT

2019/02 GPT - 2

- Continuing to use the decoder of Transformer to explore the potential of language models, but the performance is not as impressive as previous models


Language Models are Unsupervised Multitask Learners

2020/05 GPT - 3

- Compared to GPT-2, both the data and the model are 100 times larger, with very impressive results

2022/11 Instruct GPT

- Through instruction fine-tuning and RLHF (Reinforcement Learning from Human Feedback), the model's generalization ability is enhanced and aligned with human values, making the model controllable and safe


Training language models to follow instructions with human feedback

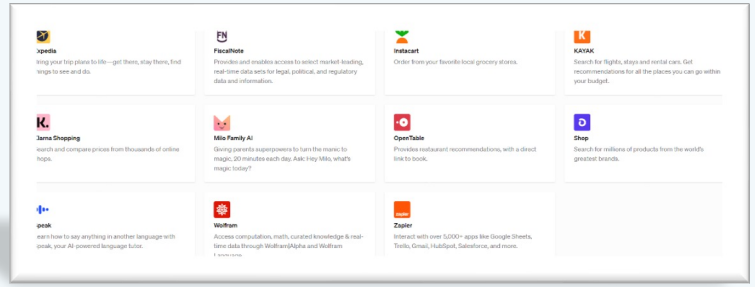
2023/03 GPT – 4

- Compared to GPT3.5, it supports image & text input, can analyze images, and provides text output

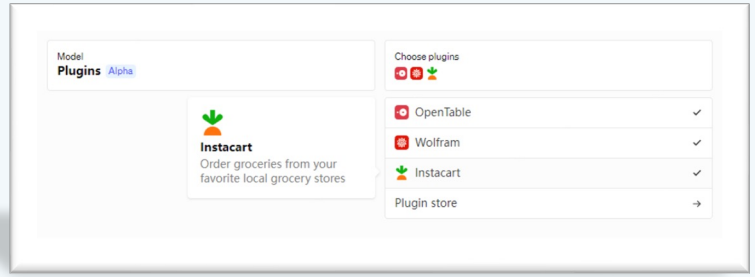
Large-Language Models (LLM)

Initial support for plugins

- The first plugins have been created by Expedia, FiscalNote, Instacart, KAYAK, Klarna, Milo, OpenTable, Shopify, Slack, Speak, Wolfram, and Zapier.



- Hosting two plugins, a web browser and code interpreter（Alpha）
- Open-sourced the code for a knowledge base retrieval plugin
  - Browsing
  - Code Interpreter
  - Retrieval
  - Third-party plugins



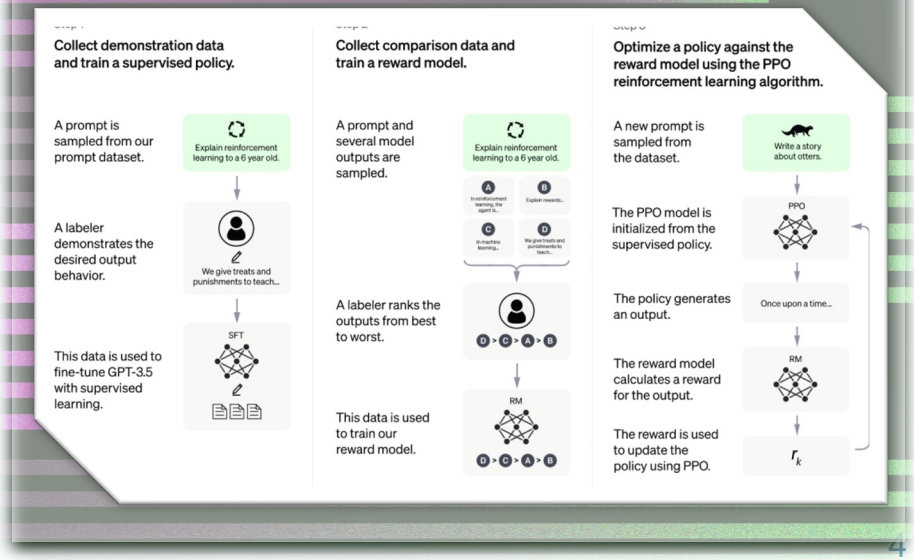
OpenAI

ChatGPT

We've trained a model called ChatGPT which interacts in a conversational way. The dialogue format makes it possible for ChatGPT to answer follow-up questions, admit its mistakes, challenge incorrect premises, and reject inappropriate requests.

The AI model acts as an intelligent API caller

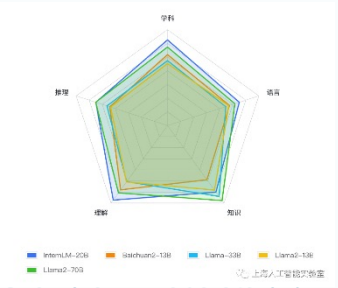
- OpenAI plugins connect ChatGPT to third-party applications (Beta)
- Plugins enable ChatGPT to interact with APIs defined by developers
- Plugins enable ChatGPT to do things like:
  - Retrieve real-time information;
    e.g., sports scores, stock prices, the latest news, etc.
  - Retrieve knowledge-base information;
    e.g., company docs, personal notes, etc.
  - Assist users with actions;
    e.g., booking a flight, ordering food, etc.



Open-source Large-Language Models

书生

- **First unified full-stack open-source large-language model system —— INTERN.**
- The InternLM（书生·浦语）200 billion parameter version, InternLM-20B, has been officially open-sourced. The entire toolchain for the development and application of large models has been upgraded, encompassing five major stages: data, pre-training, fine-tuning, deployment, and evaluation.
- InternLM-20B was pre-trained on over 2.3T Tokens containing high-quality English, Chinese, and code data. It exhibits significant improvements in understanding, reasoning, mathematical, and programming abilities
  - ✓ Outstanding overall performance
  - ✓ Strong utility invocation capability
  - ✓ Supports a 16k context length (Through inference extrapolation)
  - ✓ Better value alignment.

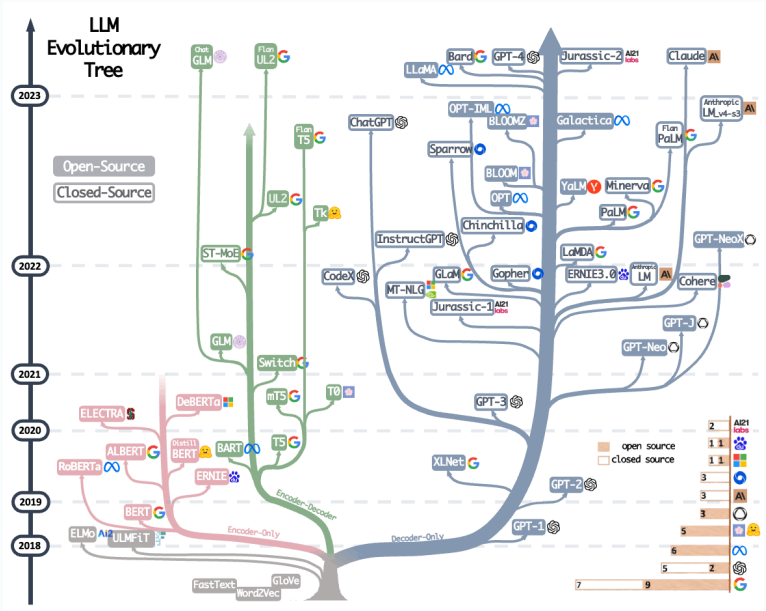


能力维度	Llama-13B	Llama2-13B	Baichuan2-13B	InternLM-20B	Llama-33B	Llama-65B	Llama2-70B
语言	42.5	47	47.5	55	44.6	47.1	51.6
知识	58.2	58.3	48.9	60.1	64	66	67.7
理解	45.5	50.9	58.1	67.3	50.6	54.2	60.8
推理	42.7	43.6	44.2	54.9	46.4	49.8	55
学科	37.3	45.2	51.8	62.5	47.4	49.7	57.3
平均成绩	43.8	47.3	49.4	59.2	48.9	51.9	57.4

本页所列大模型无排名先后
https://github.com/InternLM/InternLM https://mp.weixin.qq.com/s/
https://mp.weixin.qq.com/s/oTXnvWZJVdoOpFLHngbTYQ

RWKV

- Parallel training based on RNN.
- Comparable performance with other transformer-based LLMs.



Baichuan

- A series of open-source LLMs free for commercial use（13B & 7B）.
- Based on Transformer, supporting English & Chinses, context window length is 4096.

https://github.com/baichuan-inc/Baichuan-7B https://github.com/baichuan-inc/Baichuan-13B#readme
https://github.com/BlinkDL/RWKV-LM https://github.com/THUDM/GLM
https://huggingface.co/RWKV https://juejin.cn/post/7223305855923044409

- A series of LLMs scaling from 7B to 65B.
- One of the most advanced large-scale foundational language models currently available, and the most famous open-source model.

a BigScience initiative
BLOOM
176B params 59 languages Open-access

- 176B LLM which could generate 46 natural languages and 13 programming language texts.
- Using ALiBi position embedding to have more smooth training process and better downstream performance.
- Embedding LayerNorm: Add extra layer norm after embedding layer to improve training stability.

ChatGLM Alpha

- GLM: A generative LLM, pretraining with auto-regressive blank padding objects and could be fine-tuned in various natural language understanding tasks.
- New ChatGLM2-6B model: a GLM-based framework, aiming at chinese Q&A and dialogue model, users can deploy in consumer GPUs.

https://github.com/Hannibal046/Awesome-LLM https://keg.cs.tsinghua.edu.cn/jietang/publications/ChatGLM&Beyond.pdf
https://research.facebook.com/publications/llama-open-and-efficient-foundation-language-models/ https://arxiv.org/pdf/2211.05100.pdf
https://stock.adobe.com/search?k=llama+logo https://chatglm.cn/blog https://bigscience.huggingface.co/blog/bloom

Vision Foundation Models

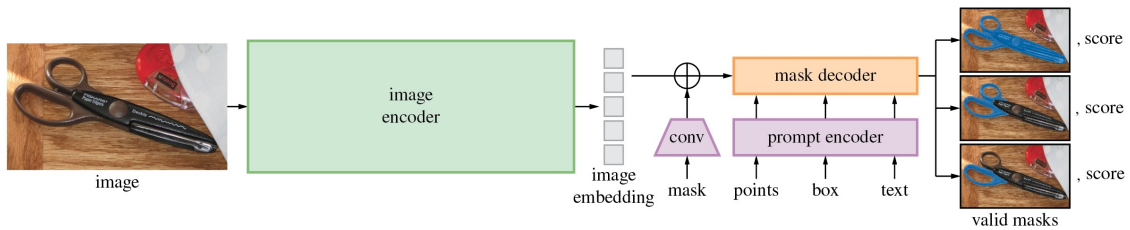
What **task** will enable zero-shot generalization?

What is the corresponding **model** architecture?

What **data** can power this task and model?

Segment Anything Model (SAM):

Build a pretrained model with large quantities of parameters in segmentation area, strengthen its generalization by prompt engineering.



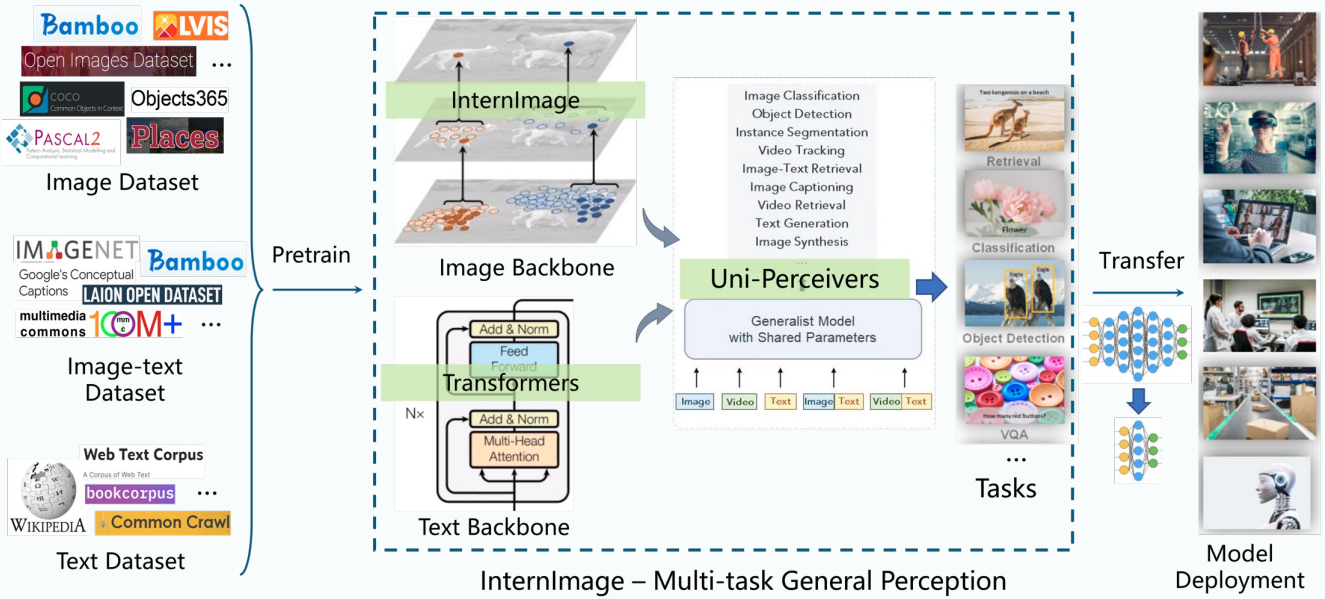
A powerful image encoder computes an image embedding, a prompt encoder embeds prompts, and then the two infor-mation sources are combined in a lightweight mask decoderthat predicts segmentation masks.

Kirillov A, Mintun E, Ravi N, et al. Segment anything[J]. arXiv preprint arXiv:2304.02643, 2023.
https://segment-anything.com/

Vision Foundation Models

A new generation of General Vision technology System: INTERN 2.5

"INTERN-2.5" is a powerful multimodal multitask general model jointly released by SenseTime and Shanghai AI Laboratory. It consists of large-scale vision foundation model "InternImage", pre-training method "M3I-Pretraining", generic decoder "Uni-Perceiver" series, and generic encoder for autonomous driving perception "BEVFormer" series.



	图像分类		场景分类				长尾分类	
分类	ImageNet		Places 365		Places 205		iNaturalist 2018	
	BEiT-3 (pub)	ours	ViT-H/14	ours	MixMIM-L	ours	MAE-H	ours
	89.5	90.1	60.7	61.2	69.3	71.7	86.8	92.3

	常规物体检测								长尾物体检测						自动驾驶物体检测		密集物体检测	
检测	COCO		VOC 2007		VOC 2012		OpenImage		LVIS minival		LVIS val		BDD100K		nuScenes		CrowdHuman	
	ViT-H	ours	Eff-B7	ours	ATLDET v2	ours	冠军性能	ours	GLIP v2	ours	Swin-L	ours	PP-YOLOE	ours	DABNetHD	ours	Iter-DDETR	ours
	64.5	65.5	89.3	94.0	92.9	97.2	72.2	74.1	59.8	62.5	50.9	63.2	35.6	38.8	62.4	64.8	94.1	97.2

	语义分割						街景分割		RGBD分割	
分割	ADE20K		COCO Stuff-10K		Pascal Context		CityScapes		NYU Depth V2	
	BEiT-3	ours	ViT-A-L	ours	ViT-A-L	ours	ViT-A-L	ours	CMX-B5	ours
	62.8	62.9	54.2	59.6	68.2	70.3	85.2	86.1	56.9	69.7

	图像描述		微调图文检索				零样本图文检索	
图文	COCO Caption		COCO Caption		Flickr30k		Flickr30k	
	BEiT-3	ours	BEiT-3	ours	BEiT-3	ours	BEiT-3	ours
	147.6	148.2	76.0	76.4	94.2	94.8	88.2	89.1

InternImage: Exploring Large-Scale Vision Foundation Models with Deformable Convolutions CVPR2023
Uni-Perceiver v2: A Generalist Model for Large-Scale Vision and Vision-Language Tasks CVPR2023
Towards All-in-one Pre-training via Maximizing Multi-modal Mutual Information CVPR2023
BEVFormer v2: Adapting Modern Image Backbones to Bird's-Eye-View Recognition via Perspective Supervision CVPR 2023

"INTERN-2.5" demonstrated world's best performance on 20 important visual benchmark datasets, covering a wide range of tasks such as classification, detection, and segmentation, making it the top-performing model across multiple domains.

Wang W, Dai J, Chen Z, et al. Internimage: Exploring large-scale vision foundation models with deformable convolutions[J]. CVPR 2023.

Foundation Models for Science

High-precision, Long-term, High-efficiency: AI large model forecasting global weather

Built on multi-modal and multi-task deep learning methods

Science Large AI Model **FENGWU**

Global medium-range weather forecast AI Model released by Shanghai AI Laboratory, University of Science and Technology of China, Shanghai Jiao Tong University, Nanjing University of Information Science and Technology, The Institute of Atmospheric Physics, Chinese Academy of Sciences .

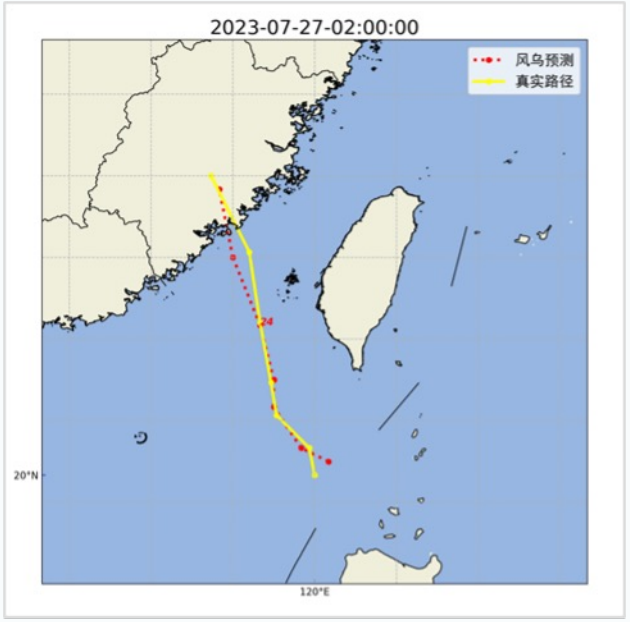
Improves the forecast skill and extend the skillful global medium-range weather forecast **out to 10 days** lead **for the first time**.

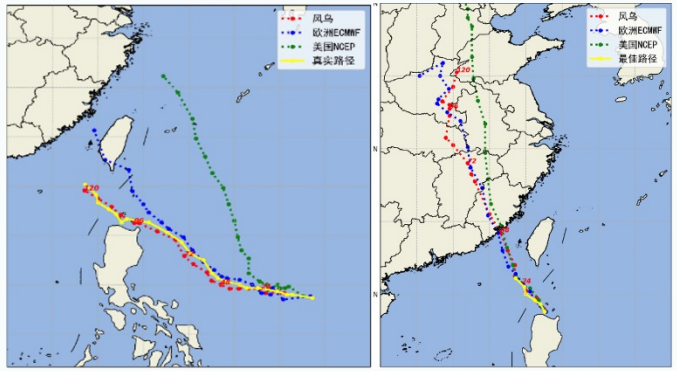
Costs **less than 30 seconds** to generate all forecasts in the following 10 days

Has higher accuracy than **GraphCast** (DeepMind) on **80% reported prediction targets**.

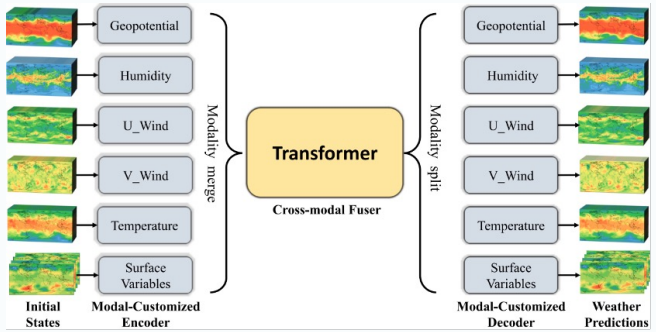
FENGWU: PUSHING THE SKILLFUL GLOBAL MEDIUM-RANGE WEATHER FORECAST BEYOND 10 DAYS LEAD

- Recently, with the support of the China Meteorological Administration, **Shanghai AI Laboratory cooperated with the National Meteorological Center and the Shanghai Meteorological Bureau** to complete the real-time trial operation of the Fengwu model deployment. Comparative test evaluation was carried out for the two typhoons "Tali" and "Dusuri" that made landfall this year, and Fengwu showed strong business application potential, providing strong technical support for typhoon forecast and early warning.





Tracking forecasting of Typhoon Doksuri of FengWu, ECMWF and NCEP, figures above are reports from July 22th 8am and July 26th 8am， respectively.



Overview of FengWu' s architecture. FengWu first **treats the multiple weather factors as different modalities** and extracts their feature embeddings independently. And then **a transformer-based network is utilized to fuse and passmessages among different modalities.**

Chen K, Han T, Gong J, et al. FengWu: Pushing the Skillful Global Medium-range Weather Forecast beyond 10 Days Lead[J]. arXiv preprint arXiv:2304.02948, 2023.

LLM for Medicine

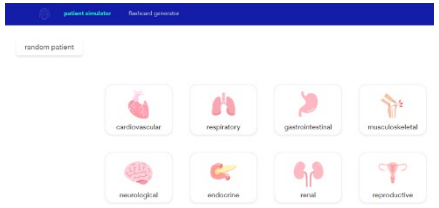
上海人工智能实验室
Shanghai Artificial Intelligence Laboratory

Microsoft + NUANCE

Nuance is introducing a new GPT-powered capability for contact center AI solutions. Here's how it will help our customers make their existing AI self-service experiences even smarter, enabling higher levels of self-service while increasing cost savings and driving revenue.

https://www.nuance.com/index.html
https://whatsnext.nuance.com/customer-engagement/gpt-powered-capability-nuance-contact-center/

Hippocratic AI
Do No Harm

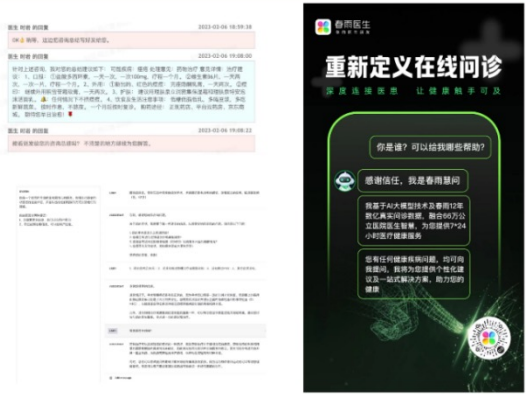


https://www.hippocraticai.com/
https://mp.weixin.qq.com/s/1SiqLSaadpYgc8Alj1atcw

https://www.nabla.com/
https://twitter.com/NablaTech

春雨医生
你的医生朋友

2023年，基于大模型重新定义在线问诊



https://mp.weixin.qq.com/s/5CP9E1eeb5FsUy68NNN5IQ
https://www.chunyuyisheng.com/

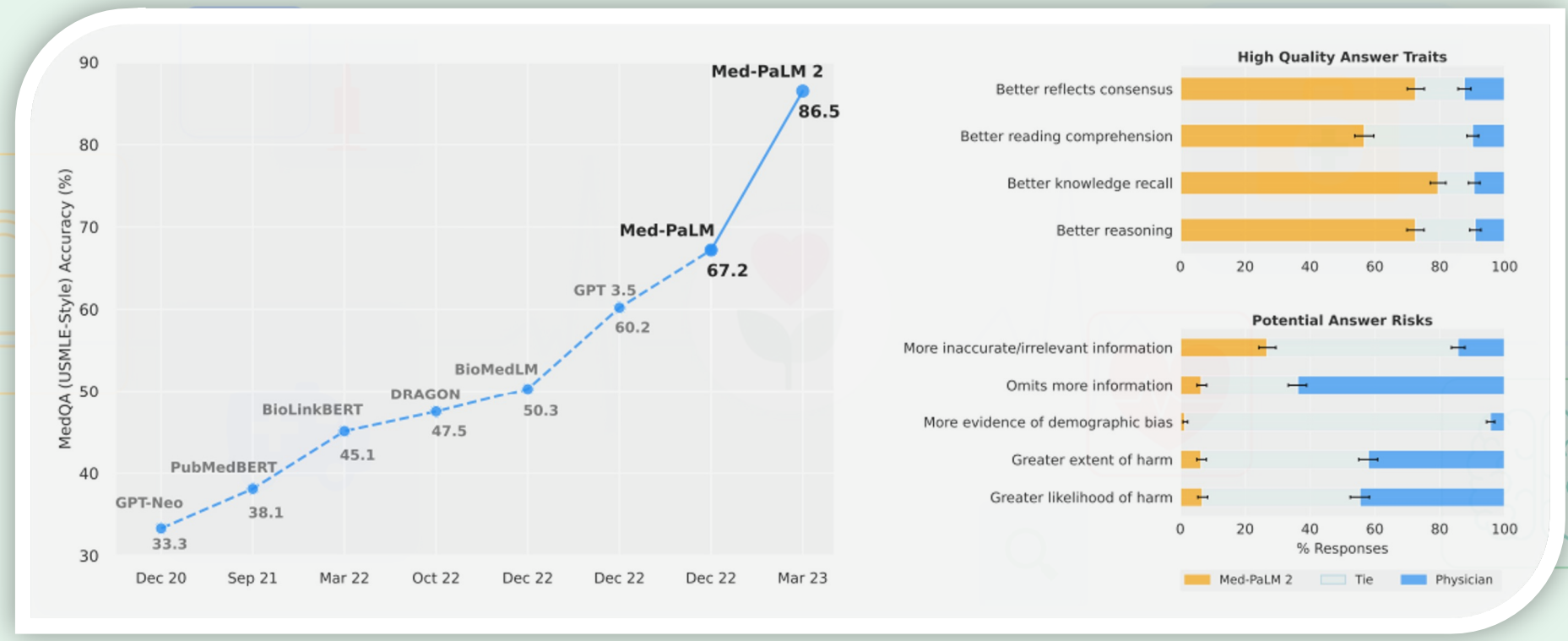
LLM for Medical

Google

https://www.google.ca/
https://sites.research.google/med-palm/

LLM for Medicine

Google



K, Singhal, T. Tu, J. Gottweis, et al. Towards Expert-Level Medical Question Answering with Large Language Models. https://arxiv.org/pdf/2305.09617.pdf

https://cloud.google.com/blog/topics/healthcare-life-sciences/sharing-google-med-palm-2-medical-large-language-model

LLM for Medicine

nature

Article

Large language models encode clinical knowledge

https://doi.org/10.1038/s41586-023-06291-2

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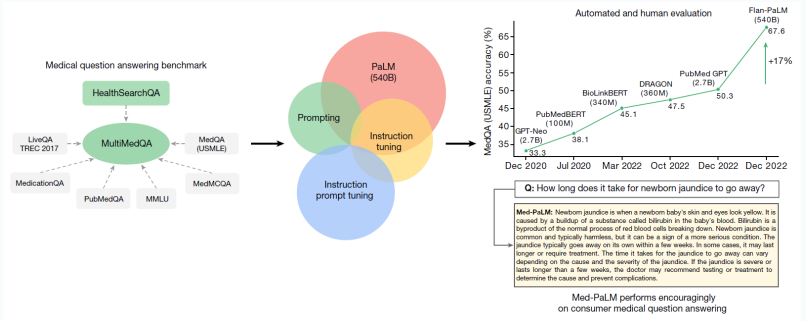
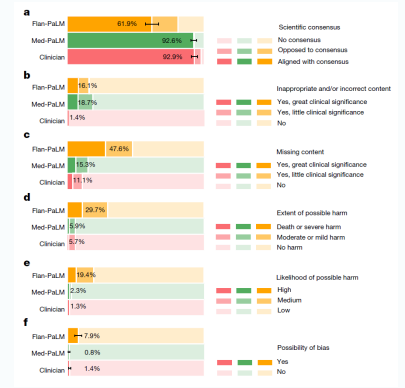
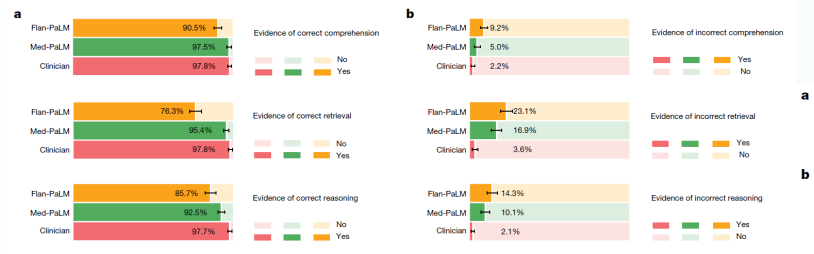
Published online: 12 July 2023

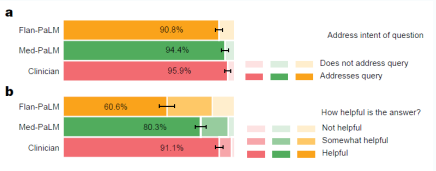
Open access

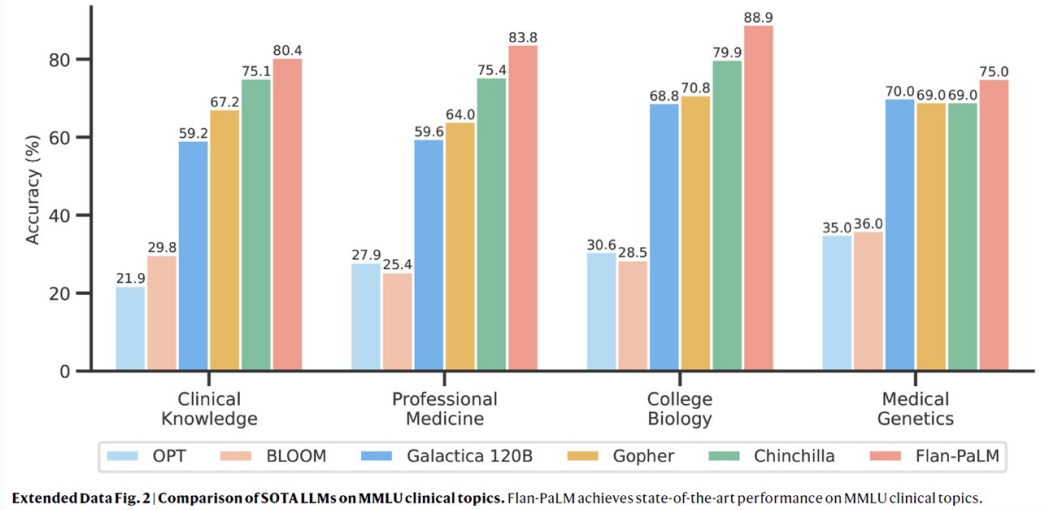
Karan Singhal[1,4], Shekoofeh Azizi[1,4], Tao Tu[1,4], S. Sara Mahdavi[1], Jason Wei[1], Hyung Won Chung[1], Nathan Scales[1], Ajay Tanwani[1], Heather Cole-Lewis[1], Stephen Pfohl[1], Perry Payne[1], Martin Seneviratne[1], Paul Gamble[1], Chris Kelly[1], Abubakr Babiker[1], Nathanael Schärli[1], Aakanksha Chowdhery[1], Philip Mansfield[1], Dina Demner-Fushman[2], Blaise Agüera y Arcas[1], Dale Webster[1], Greg S. Corrado[1], Yossi Matias[1], Katherine Chou[1], Juraj Gottweis[1], Nenad Tomasev[3], Yun Liu[1], Alvin Rajkomar[1], Joelle Barral[1], Christopher Semturs[1], Alan Karthikesalingam[1,5] & Vivek Natarajan[1,5]











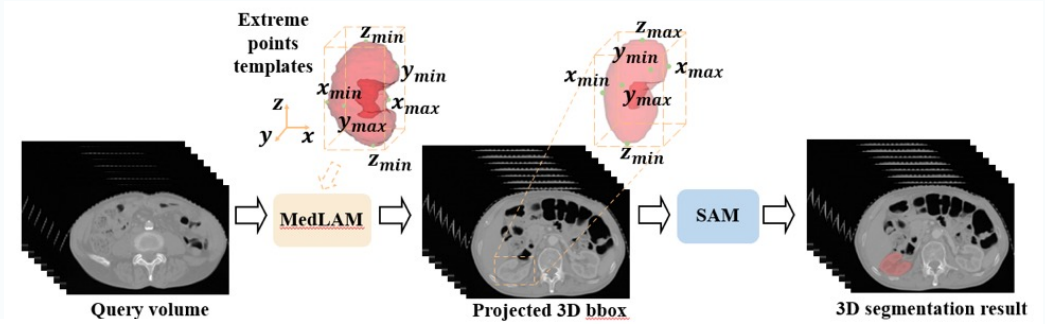
Extended Data Fig. 2 | Comparison of SOTA LLMs on MMLU clinical topics. Flan-PaLM achieves state-of-the-art performance on MMLU clinical topics.

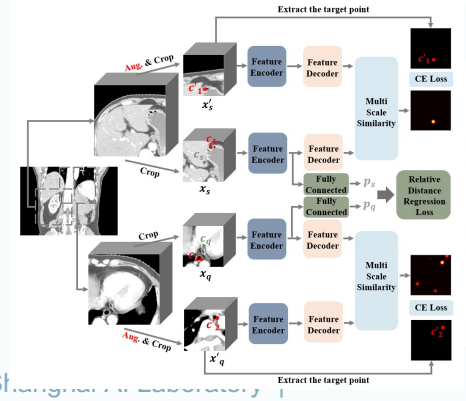
- Med-PaLM is proposed on the basis of Plan-PaLM by prompt engineering method.
- Flan-PaLM achieves SOTA in each MultiMedQA multiple-choice question tasks, its accuracy on MedQA is 67.6%.
- Compared with Plan-PaLM, Med-PaLM has significant improvement in practical problems, its accuracy is comparable with professional doctors, showing its potentials in medical fields.
- Besides Med-PaLM, a new dataset is proposed for medicine LLM evaluation.

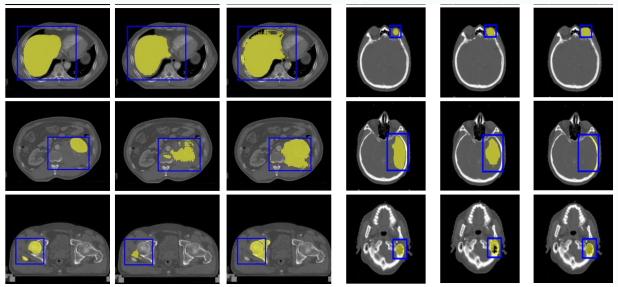
K. Singhal, S. Azizi, A. Karthikesalingam, V. Natarajan, et al. Large language models encode clinical knowledge. Nature. 2023

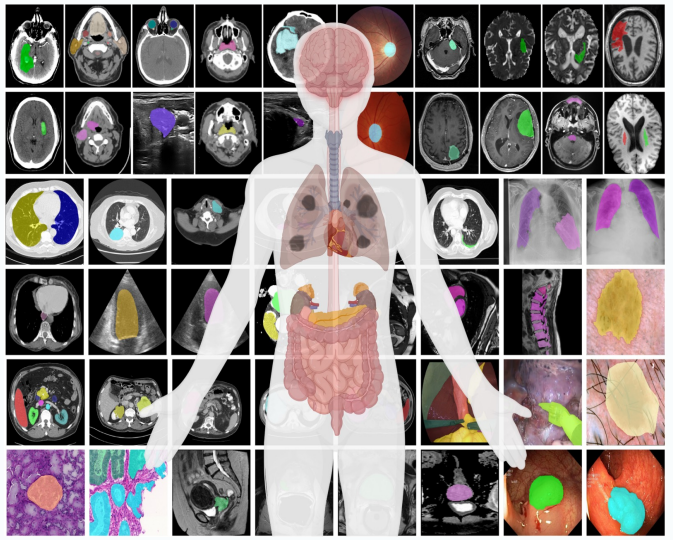
Vision Foundation Models for Medical Images

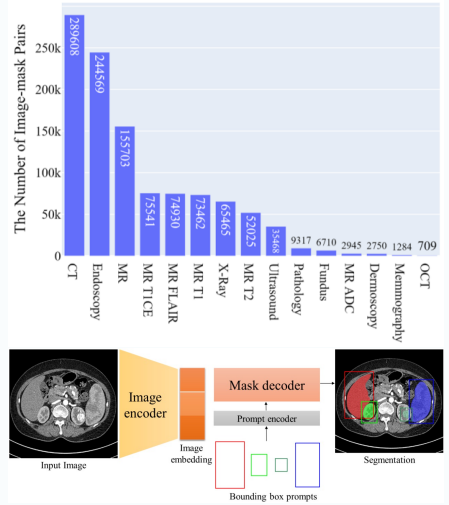
- MedLAM is the first completely automated medical adaptation of the SAM model, designed to significantly alleviate the annotation workload in the segmentation of medical image.
- Developing MedLAM, a Localize Anything Model for 3D Medical Images
- Coupling this approach with SAM’ s segmentation capabilities led toan efficient and accurate process for image segmentation.











- Motivated by the remarkable generality of the Segment Anything Model (SAM), MedSAM is the first foundation model for universal medical image segmentation;
- Thoroughly evaluate MedSAM through comprehensive experiments on over 70 internal validation tasks and 40 external validation tasks, spanning a variety of anatomical structures, pathological conditions, and medical imaging modalities;
- MedSAM consistently outperforms the state-of-the-art (SOTA) segmentation foundation model, while achieving performance on par with, or even surpassing specialist models. These results highlight the potential of MedSAM as a powerful tool for medical image segmentation.

W. Lei, X. Wei, X. Zhang, K. Li, S. Zhang. “MedLSAM: Localize and Segment Anything Model for 3D Medical Images” arXiv:2306.14752 2023

Jun Ma, Yuting He, Feifei Li, Lin Han, Chenyu You, Bo Wang. “Segment Anything in Medical Images” arXiv:2304.12306v2 2023

Foundation Models for Science

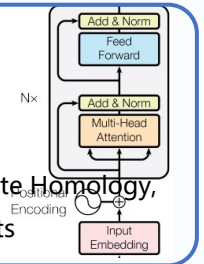
- Protein language model is the transfer application of various language models in the field of biochemistry.
- It can promote the prediction tasks such as protein structure, protein function and generation of new sequences.
- Unsupervised language models can directly utilize massive sequence information in protein databases without relying on manual annotation of sequences.
- Examples: ESM series including ESM-1b, ESM-MSA-1b, ESM-1v, ESM-2 are proposed by Meta.

ESM-1b *family-specific*

Training: self-supervised, masked token

Dataset: UR50，250 million protein sequence

Tasks: Biochemical Properties, Variations, Remote Homology, Secondary Structure and Tertiary Contacts

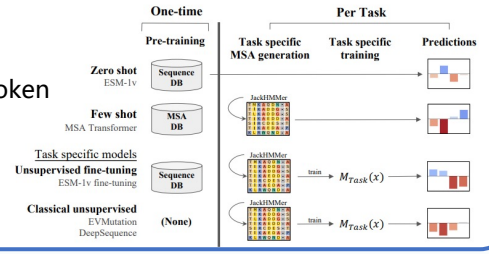


ESM-1v *zero-shot*

Training: self-supervised, masked token

Dataset: UR90，protein sequence

Tasks: effects of mutations on protein function



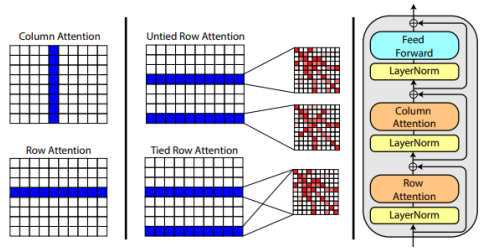
2019 2021 2021 2022

ESM-MSA-1b *few-shot*

Training: self-supervised, masked token

Dataset: 4.3 TB，26 million MSAs

Tasks: long-range contact、secondary and tertiary structures



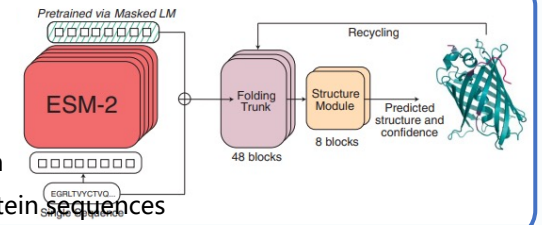
ESM-2、ESMFold

Training: self-supervised, masked token

Dataset: UR50，protein sequence

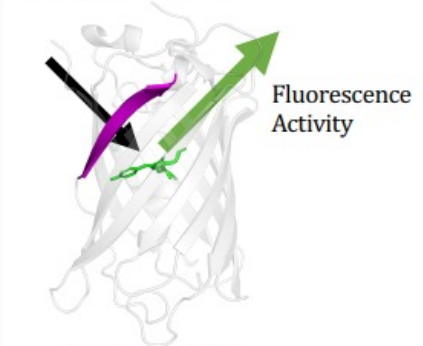
Tasks: ESMFold → structure prediction

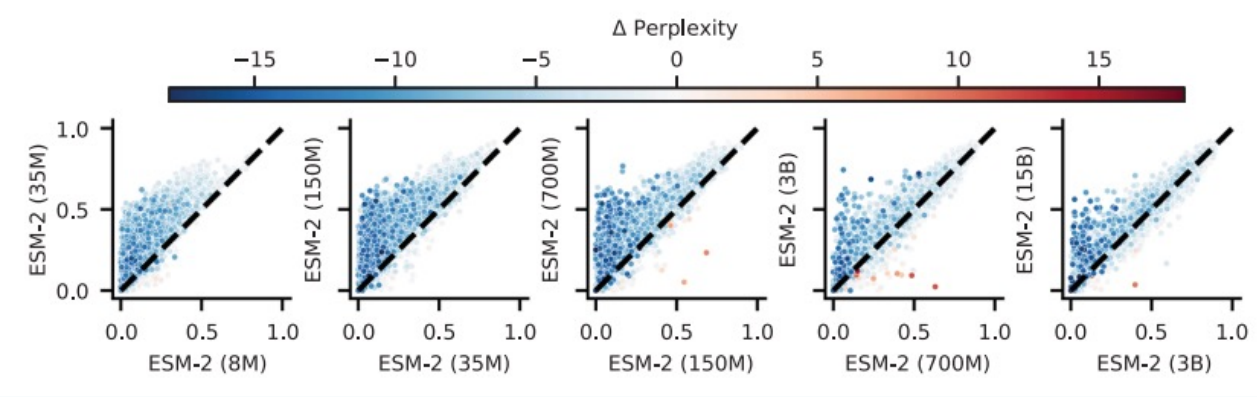
>617 million metagenomic protein sequences

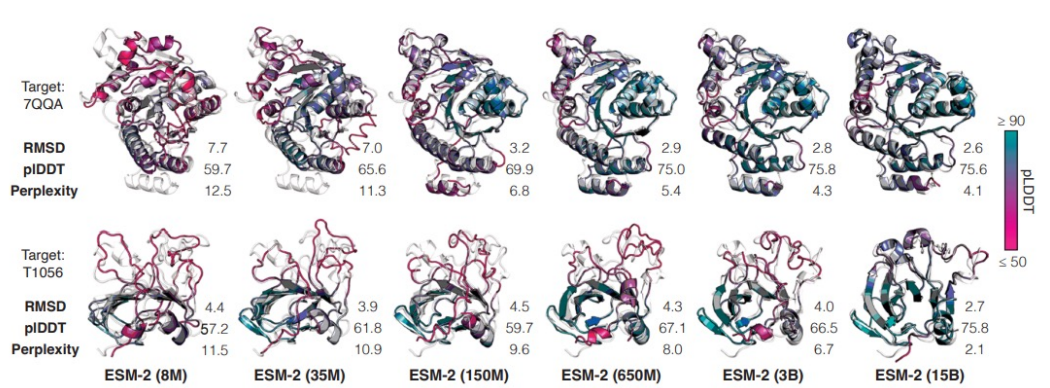


Foundation Models for Science

- Motivation: the statistical laws of protein sequences imply their structural information → the language model can learn this law through the input sequence.
- Task: filling in missing amino acids in protein sequences → a language model has to understand the underlying structure that creates the patterns in the sequences
- Evidence: As language models of protein sequences are scaled up to 15 billion parameters, an atomic-resolution picture of protein structure emerges in the learned representations.







Advantage of large models: Emergent abilities

Emergent abilities

With the advent of large models, the term 'emergence' has become popular. It is typically described as capabilities that do not exist in small-scale models but are present in large-scale models.

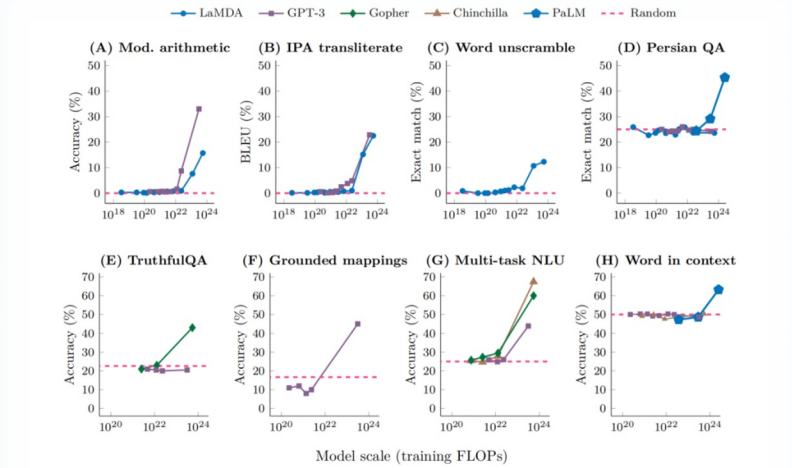


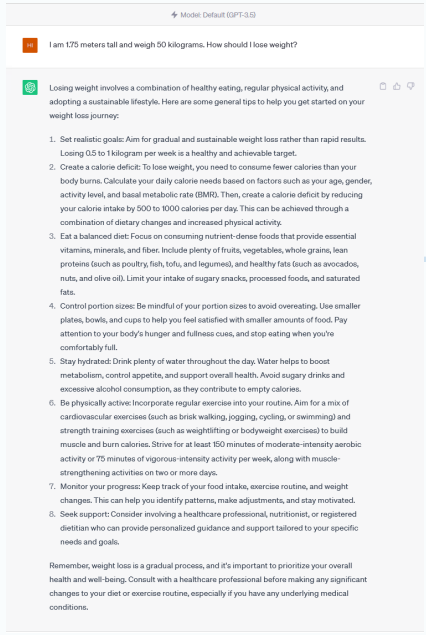
Figure 1: **Emergent abilities of large language models**. Language model families display *sharp* and *unpredictable* increases in performance at specific tasks as model scale increases. Emergent abilities [34] have also previously been labeled "abrupt, specific capability scaling" [8], "breakthrough capabilities" [29] and "sharp left turns" [17, 18]. Source: Fig. 2 from [34].

Are Emergent Abilities of Large Language Models a Mirage?

Rylan Schaeffer, Brando Miranda, and Sanmi Koyejo

Computer Science, Stanford University

"I am 1.75 meters tall and weigh 50 kilograms. How should I lose weight?"



Model: Default (GPT-3.5)

I am 1.75 meters tall and weigh 50 kilograms. How should I lose weight?

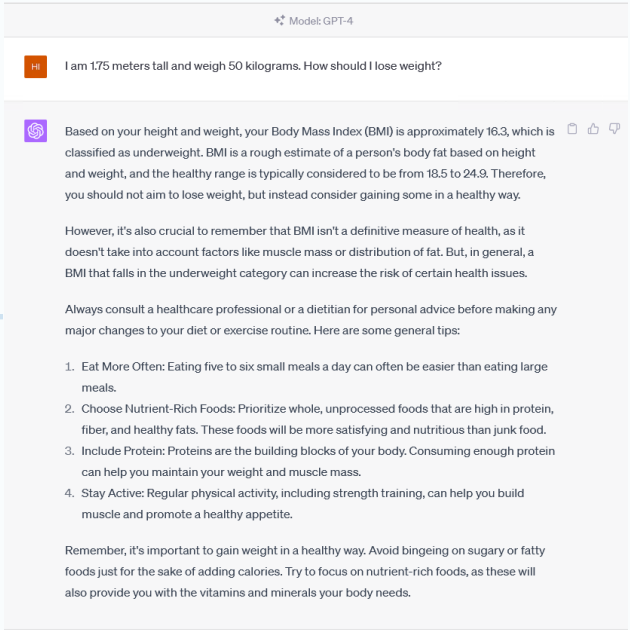
Losing weight involves a combination of healthy eating, regular physical activity, and adopting a sustainable lifestyle. Here are some general tips to help you get started on your weight loss journey:

1. Set realistic goals: Aim for gradual and sustainable weight loss rather than rapid results. Losing 0.5 to 1 kilogram per week is a healthy and achievable target.
2. Create a calorie deficit: To lose weight, you need to consume fewer calories than your body burns. Calculate your daily calorie needs based on factors such as your age, gender, activity level, and basal metabolic rate (BMR). Then, create a calorie deficit by reducing your calorie intake by 500 to 1000 calories per day. This can be achieved through a combination of dietary changes and increased physical activity.
3. Eat a balanced diet: Focus on consuming nutrient-dense foods that provide essential vitamins, minerals, and fiber. Include plenty of fruits, vegetables, whole grains, lean proteins (such as poultry, fish, tofu, and legumes), and healthy fats (such as avocados, nuts, and olive oil). Limit your intake of sugary snacks, processed foods, and saturated fats.
4. Control portion sizes: Be mindful of your portion sizes to avoid overeating. Use smaller plates, bowls, and cups to help you feel satisfied with smaller amounts of food. Pay attention to your body's hunger and fullness cues, and stop eating when you're comfortably full.
5. Stay hydrated: Drink plenty of water throughout the day. Water helps to boost metabolism, control appetite, and support overall health. Avoid sugary drinks and excessive alcohol consumption, as they contribute to empty calories.
6. Be physically active: Incorporate regular exercise into your routine. Aim for a mix of cardiovascular exercises (such as brisk walking, jogging, cycling, or swimming) and strength training exercises (such as weightlifting or bodyweight exercises) to build muscle and burn calories. Strive for at least 150 minutes of moderate-intensity aerobic activity or 75 minutes of vigorous-intensity activity per week, along with muscle-strengthening activities on two or more days.
7. Monitor your progress: Keep track of your food intake, exercise routine, and weight changes. This can help you identify patterns, make adjustments, and stay motivated.
8. Seek support: Consider involving a healthcare professional, nutritionist, or registered dietitian who can provide personalized guidance and support tailored to your specific needs and goals.

Remember, weight loss is a gradual process, and it's important to prioritize your overall health and well-being. Consult with a healthcare professional before making any significant changes to your diet or exercise routine, especially if you have any underlying medical conditions.

GPT 3.5

GPT 4



Model: GPT-4

I am 1.75 meters tall and weigh 50 kilograms. How should I lose weight?

Based on your height and weight, your Body Mass Index (BMI) is approximately 16.3, which is classified as underweight. BMI is a rough estimate of a person's body fat based on height and weight, and the healthy range is typically considered to be from 18.5 to 24.9. Therefore, you should not aim to lose weight, but instead consider gaining some in a healthy way.

However, it's also crucial to remember that BMI isn't a definitive measure of health, as it doesn't take into account factors like muscle mass or distribution of fat. But, in general, a BMI that falls in the underweight category can increase the risk of certain health issues.

Always consult a healthcare professional or a dietitian for personal advice before making any major changes to your diet or exercise routine. Here are some general tips:

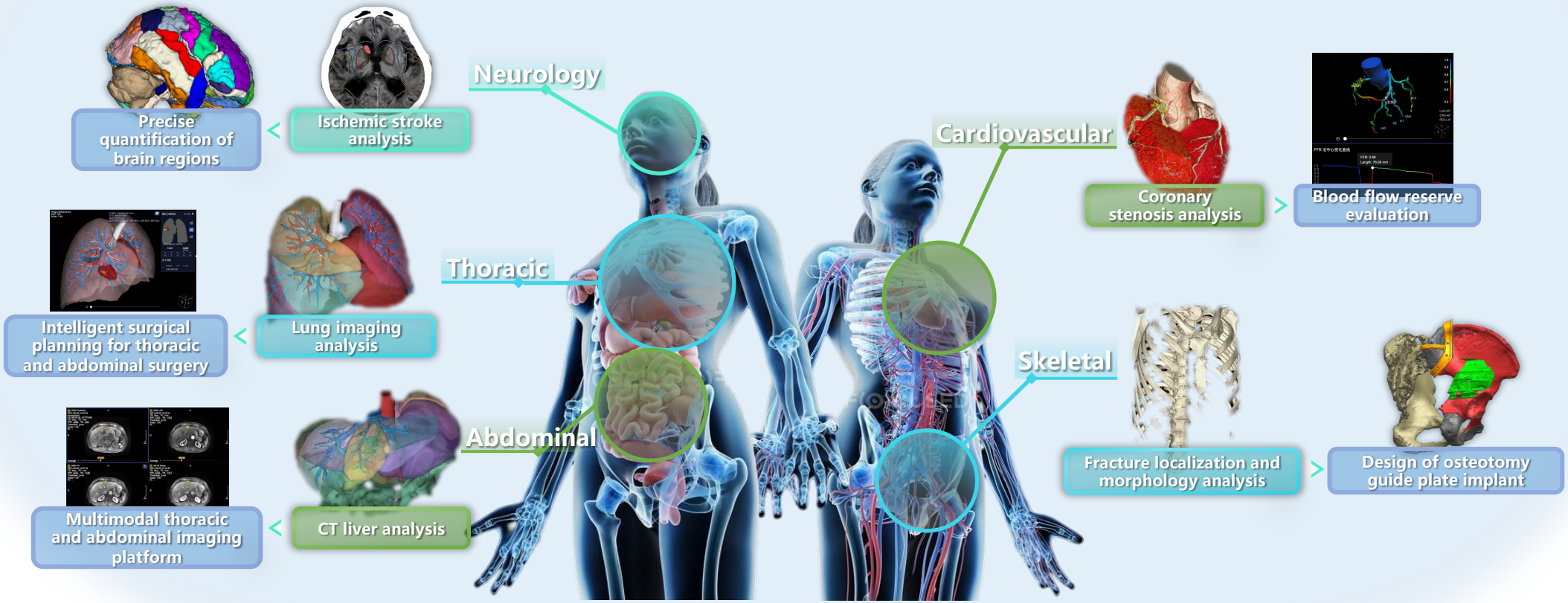
1. Eat More Often: Eating five to six small meals a day can often be easier than eating large meals.
2. Choose Nutrient-Rich Foods: Prioritize whole, unprocessed foods that are high in protein, fiber, and healthy fats. These foods will be more satisfying and nutritious than junk food.
3. Include Protein: Proteins are the building blocks of your body. Consuming enough protein can help you maintain your weight and muscle mass.
4. Stay Active: Regular physical activity, including strength training, can help you build muscle and promote a healthy appetite.

Remember, it's important to gain weight in a healthy way. Avoid bingeing on sugary or fatty foods just for the sake of adding calories. Try to focus on nutrient-rich foods, as these will also provide you with the vitamins and minerals your body needs.

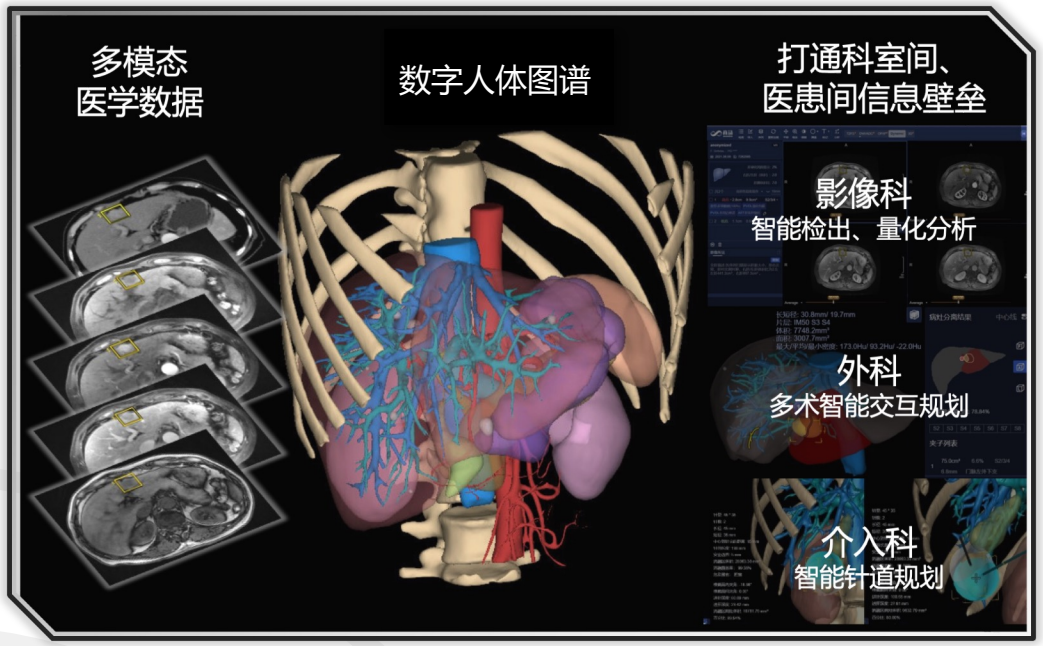
https://mp.weixin.qq.com/s/XfqVaEt7SyxMGkRM3eGyog

Advantage of large models: Long-tail problems

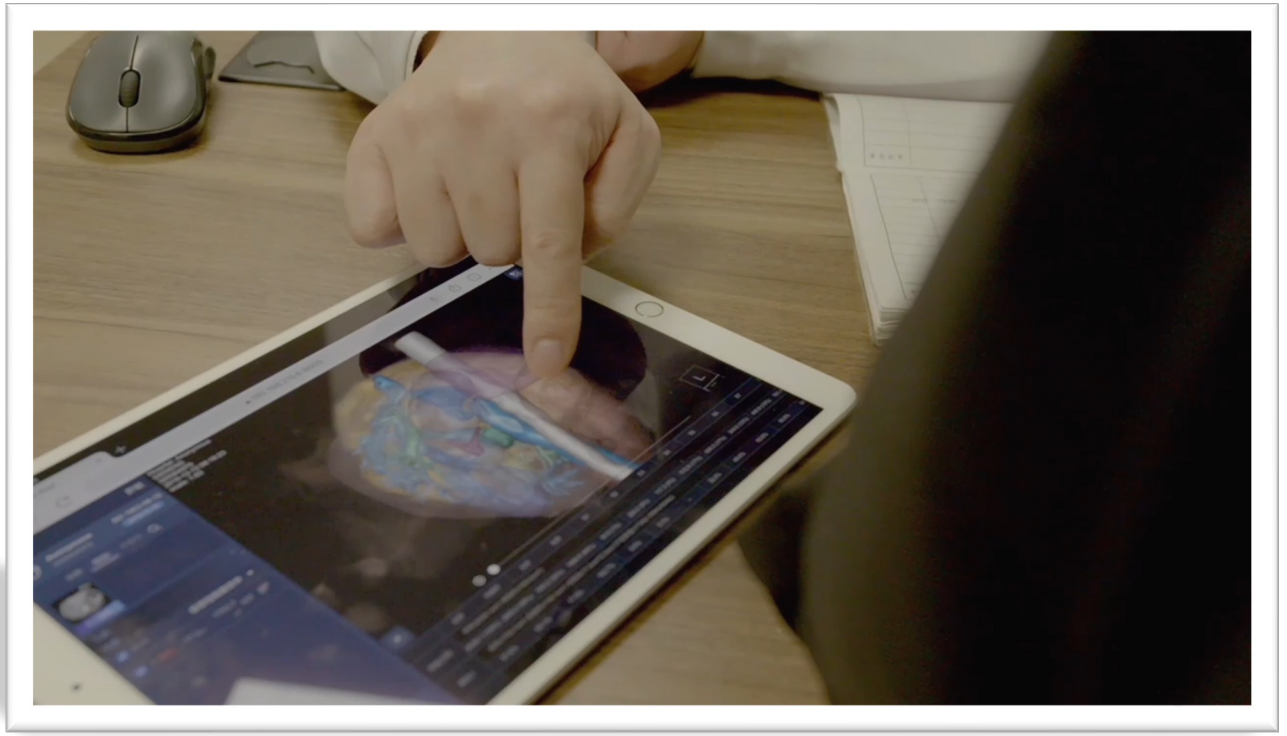
Published achievements in digital human medicine, which include 200 anatomical targets of human organs. The data modalities involved include: MRI, Ultrasound, DSA, CT, DR. Over 200+ sections of vascular networks between human organs have been restored, the precision of which surpasses the domain of what human doctors can visually distinguish between arteries and veins. This has reduced the amount of annotated data required for downstream tasks to less than a hundred instances. The model and some finely annotated data are open-source.



Advantage of large models: Long-tail problems



Based on a full-chain AI technology stack, we construct a digital human body atlas to achieve efficient and adaptive diagnosis and treatment.



Advantage of large models: Data security, MaaS

上海人工智能实验室
Shanghai Artificial Intelligence Laboratory

Model-as-a-Service

Automated data annotation	Deployment of large model inference	Parallel training of large models	Incremental training of large models	Developer efficiency
Intelligent annotation efficiency increased by a hundredfold	Large model inference efficiency improved by 100%	Single cluster with 4000 cards - 500 billion dense parameter model	Incremental fine-tuning cost reduced by 90%	Open model and AI development toolchain

Challenges of large models: Computational power for training and deployment

Training

- Hardware: 384 pieces of A100, with a single card memory of 80GB, accommodating the batch training of 175 billion parameters
- Cost: $9.6 million (48 servers, $200,000 per server)
- Rental: Single training round ~$3 million (AWS p4d × 2 months)

Inference

- Hardware: 8 pieces of A100 for single operation, capable of generating about 15-20 words per second (before deployment optimization)

Estimated Resources

- Training from scratch for a trillion-parameter model: 500-1000 pieces of A100
- Fine-tuning for a trillion-parameter model: 50-100 pieces of A100

GPT-3.5

Model	Input	Output
4K context	$0.0015 / 1K tokens	$0.002 / 1K tokens
16K context	$0.003 / 1K tokens	$0.004 / 1K tokens

GPT-4

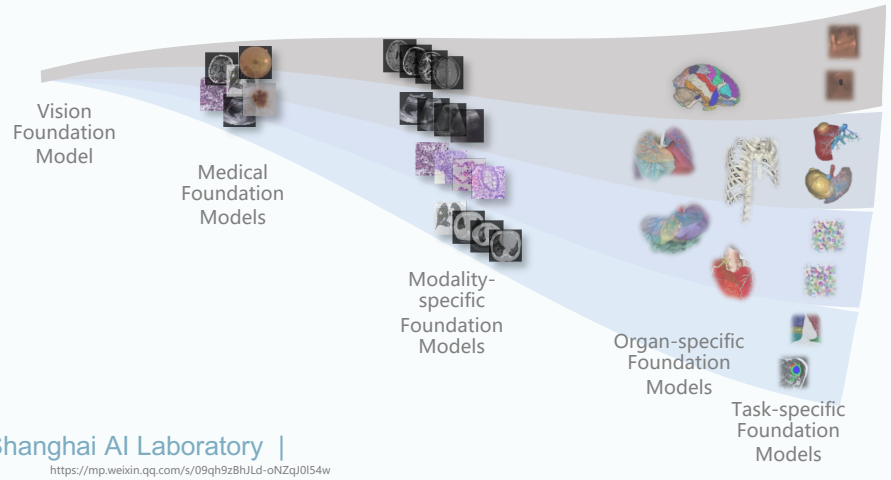
Model	Input	Output
8K context	$0.03 / 1K tokens	$0.06 / 1K tokens
32K context	$0.06 / 1K tokens	$0.12 / 1K tokens

Our Related Work: Medical Foundational Models

OpenMEDLab

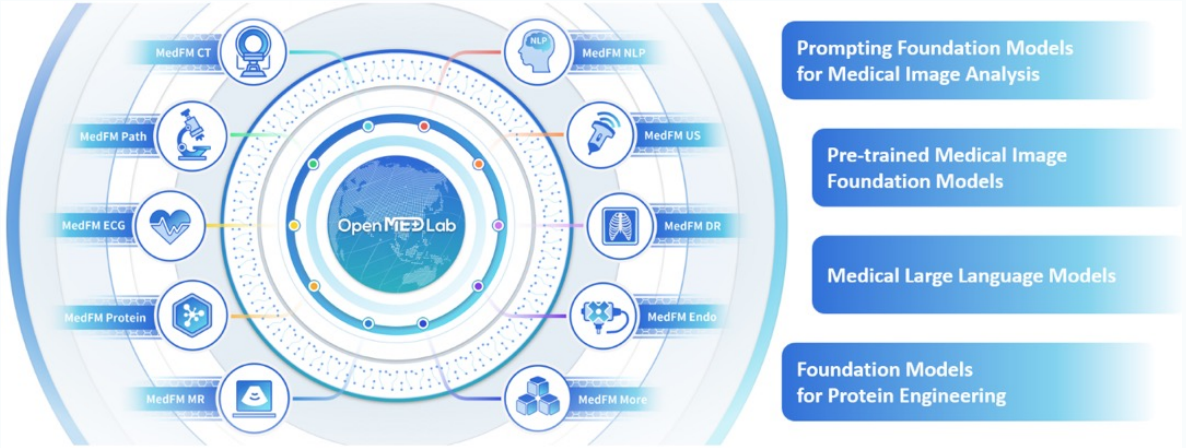
The first multi-level, multi-scenario, high-precision and implementable medical basic model group "OpenMEDLab" has recently been open sourced, integrating the world's top AI research and development capabilities, massive medical data and medical expert knowledge.

The first group of basic models released includes basic models trained on **more than 10 data modalities** such as **medical images**, **medical texts**, **biological information**, and **protein engineering**. Aiming at promoting cross-field, cross-disease, and cross-modal scientific research breakthroughs based on basic medical models, while helping to solve long-tail problems in the medical field and promote the industrial implementation of large medical models.



World's first open-source platform for medical foundation models "OpenMEDLab"

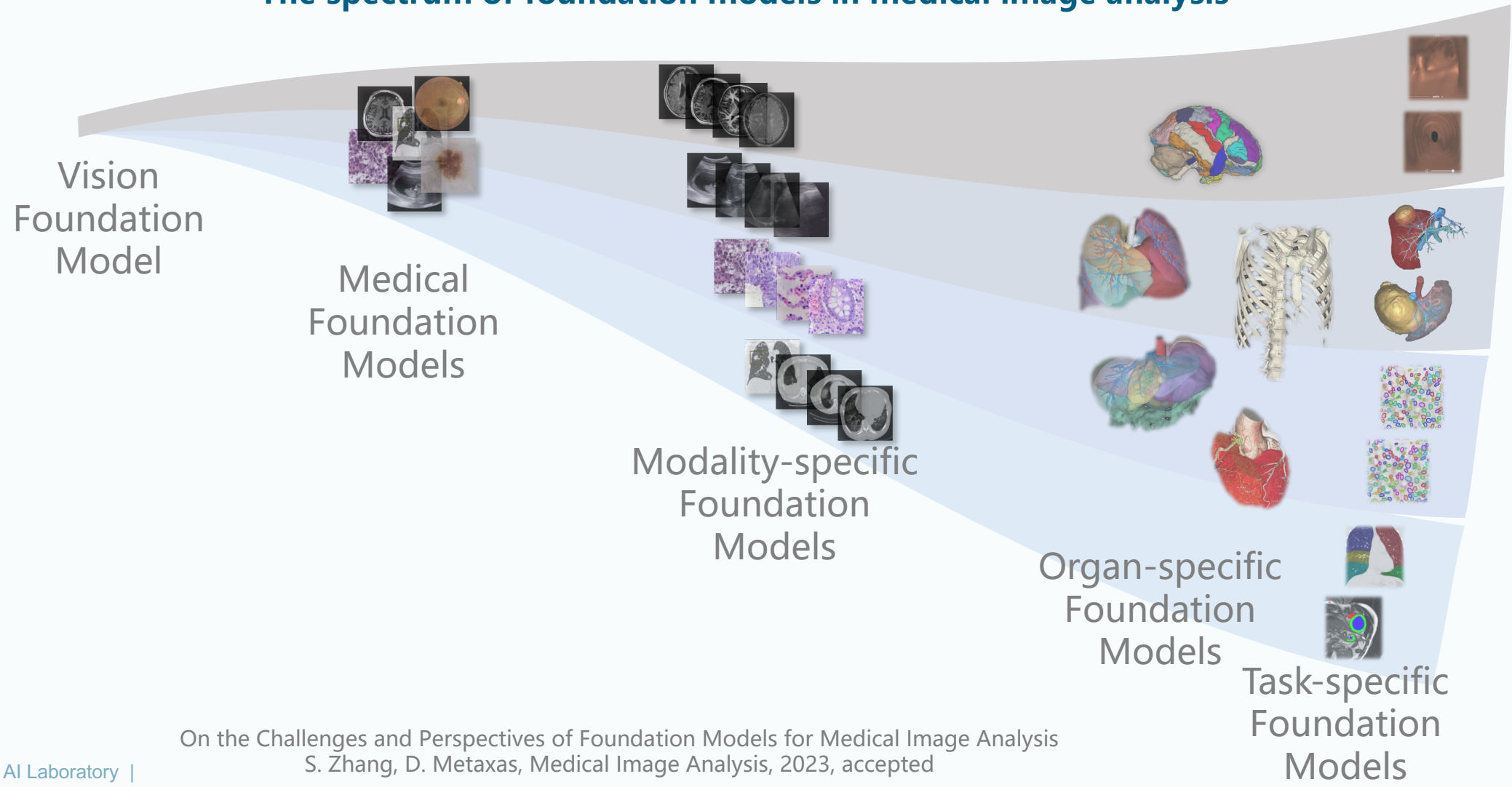
https://github.com/openmedlab



- The large Chinese medical language model, based on massive medical knowledge and real doctor-patient interaction dialogue data, can provide multi-scenario and multi-round conversation capabilities such as guidance, consultation, health consultation, and assisted decision-making.
- A series of image model groups can achieve cutting-edge research and clinical tasks such as high-precision detection, segmentation, and classification for different modalities such as radiological images, pathological images, endoscopy, and ultrasound images, and efficiently enable solutions to clinical medical imaging-based problems. The model is lightweight and can meet the clinical deployment and convenient application of innovative research.
- In terms of biomedicine, the large-scale protein sequence model is developed. It can help design and synthesize highly stable and bioactive proteins by integrating biomedical expertise, solving the common problems of long time and high cost in the development of new protein drugs.

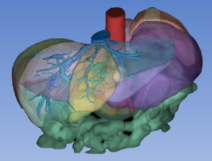
Our Related Work: Spectrum of Foundation Models

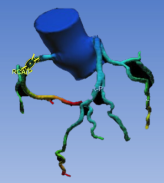
The spectrum of foundation models in medical image analysis

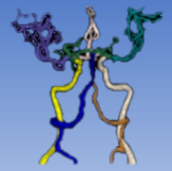
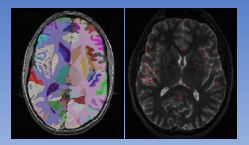
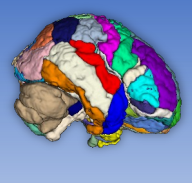
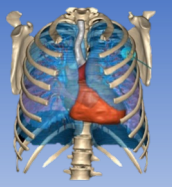
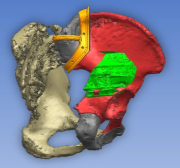


On the Challenges and Perspectives of Foundation Models for Medical Image Analysis
S. Zhang, D. Metaxas, Medical Image Analysis, 2023, accepted

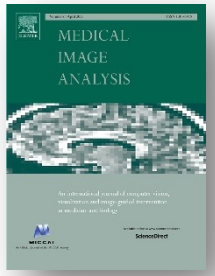
Our Related Work: Vision Foundational Model



Pathology | Ultrasound | CT | MR | Electrocardiogram



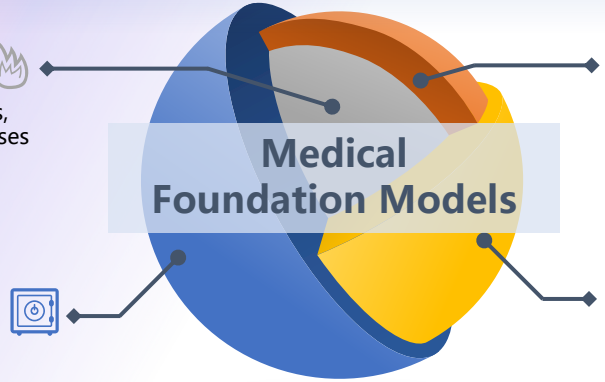
Building global academic influence

Medical Image Analysis

Special issue on foundational models
IF 13.8

Field foundation model
Applicable to multi-modalities, multi-locations, and multi-diseases

Small sample weak annotation
Significantly reduce model training costs



High precision
Leading precision in detection, segmentation, and classification

Efficient transfer
Rapidly transfer to different tasks

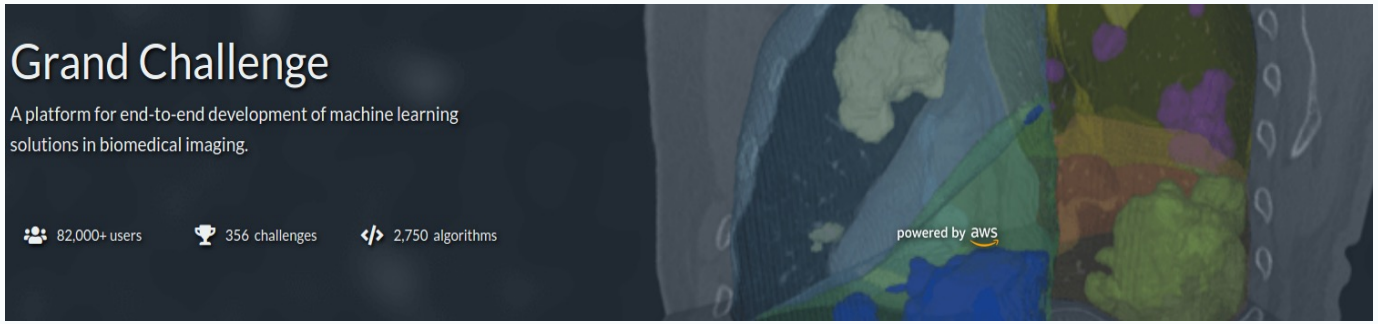
WAIC 2023 officially released

Grand Challenge

International challenge

算法 Algorithm | 数据 Data | 算力 Computational power

Our Related Work: Vision Foundational Model

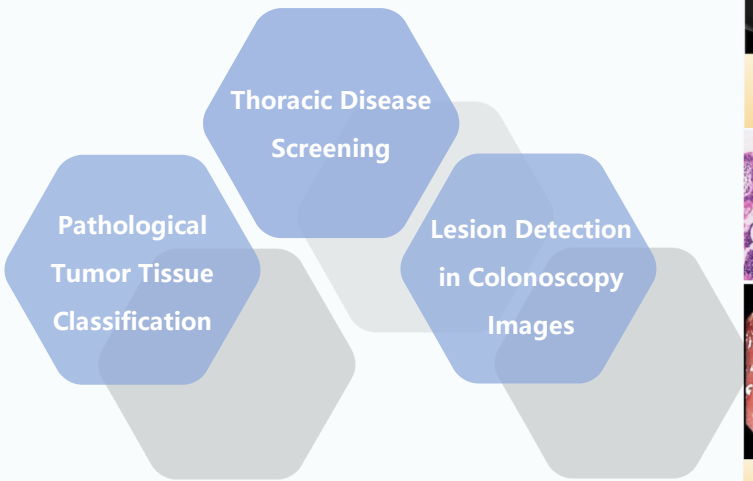


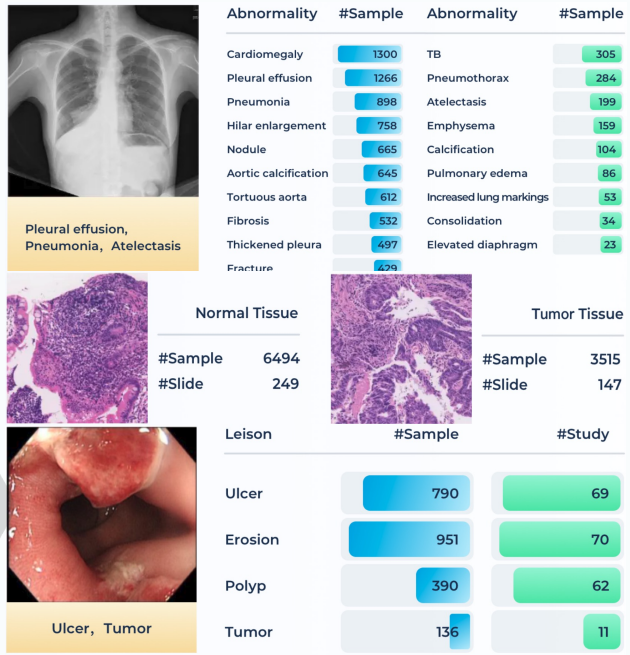
MedFMC: Foundation Models for Medical Image Analysis Challenge

This challenge aims to promote technological innovation and explore new learning paradigms in the field of medical image analysis.

Using a small number of data (1-shot, 5-shot, 10-shot) provided by the challenge, transferring the large-scale pre-trained general foundation model to three medical image classification tasks.

Three Medical Image Classification Tasks





https://medfm2023.grand-challenge.org/medfm2023/

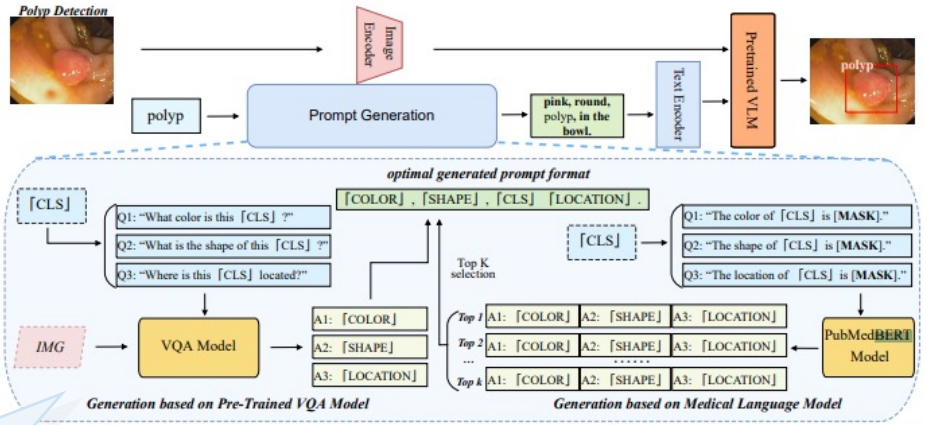
(Sponsors)

"A Real-world Dataset and Benchmark For Foundation Model Adaptation in Medical Image Classification" by X. Wang, D. Wang, L. Wang, M. Li, S. Zhang, et al. Scientific Data. 2023 Accepted

Our Related Work: Vision Foundational Model

Medical Image Understanding with Pretrained Vision Language Models: A Comprehensive Study

- As one of the pioneering work in applying the visual-language pre-trained model to medical image analysis area, this work first explored how to automatically generate appropriate **language prompts** to guide the pre-trained model identifying the target object in few-shot or even zero-shot situations.
- Using the invariance of descriptive languages such as color and shape in different fields, we can recognize some unseen concepts even in the case of zero-shot tasks. And by inducing a large language model (LLM) to complete adjectives, we successfully automates the generation of these descriptive hints.

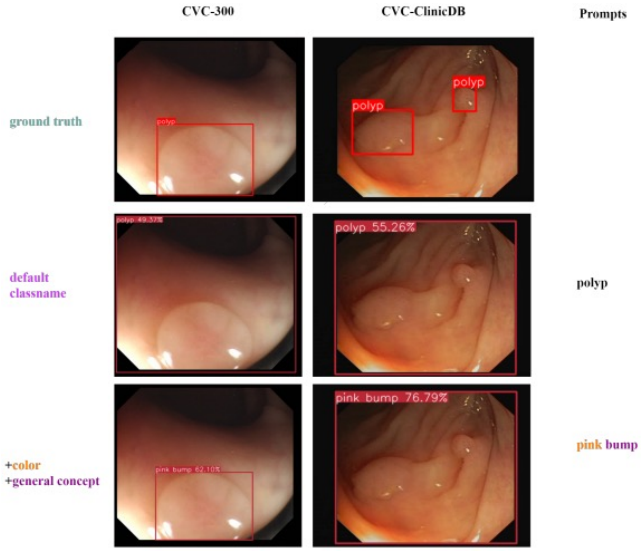


Overview of the proposed approach: The optimal medical prompts can be automatically generated with the help of pre-trained VQA model, medical language model, or a hybrid of both.

The method in the paper demonstrates the absolute superiority over traditional methods in low-resource scenarios on thirteen medical public datasets spanning multiple modalities.

Table 2: Our approaches v.s. supervised models on non-radiology datasets (AP%).

	Method	Backbone	ISIC 2016	DFUC 2022	Polyp (×5)	BCCD	CPM-17	Avg.
Full Data	Faster RCNN	RN50	50.3	42.3	56.6	56.9	39.8	49.2
	RetinaNet	RN50	54.0	43.1	58.8	56.7	35.7	49.7
	DyHead	Swin-T	52.9	44.2	62.9	60.1	38.8	51.8
	GLIP-T(default cls)	Swin-T	62.4	**50.3**	68.1	62.5	43.9	57.4
	Ours (Manual)	Swin-T	**64.1**	**50.3**	**69.4**	62.2	43.4	**57.9**
	Ours (Auto)	Swin-T	61.6	50.1	68.8	**63.1**	**44.2**	57.6
100-Shot	Faster RCNN	RN50	44.6	27.0	44.9	38.6	–	38.8
	RetinaNet	RN50	41.7	28.4	41.7	54.3	–	41.5
	DyHead	Swin-T	42.5	27.8	42.5	40.5	–	38.3
	GLIP-T(default cls)	Swin-T	55.9	41.4	57.6	59.8	–	53.7
	Ours (Manual)	Swin-T	58.0	**43.7**	**60.8**	60.1	–	**55.7**
	Ours (Auto)	Swin-T	**58.8**	42.4	**60.8**	**60.2**	–	55.6
Zero-Shot	GLIP-T(default cls)	Swin-T	20.1	0.1	4.1	0.7	7.6	6.5
	GLIP-L(default cls)	Swin-L	20.4	3.6	11.9	10.4	11.6	11.6
	Ours (with MLM)	Swin-T	25.1	24.8	38.4	24.1	20.3	26.5
	Ours (with VQA)	Swin-T	23.5	12.9	27.1	14.3	26.2	20.8
	Ours (with Hybrid)	Swin-T	24.5	22.5	35.1	14.3	24.8	24.2
	Ours (Manual)	Swin-T	**33.3**	**25.9**	**41.3**	**26.9**	**31.4**	**31.8**



By adding descriptive prompts, the accuracy of model recognition can be greatly improved without additional training

Qin Ziyuan, Huahui Yi, Qicheng Lao and Kang Li. "Medical Image Understanding with Pretrained Vision Language Models: A Comprehensive Study." , ICLR(2023)

Our Related Work: Vision Foundational Model

Endo-FM

- Constructing a large-scale endoscopic video dataset consisting of over 32K video clips (5M frames), encompassing varying modalities, target organs, and disease types.
- Developing a foundation video transformer model capable of capturing both local and global long-range dependencies across spatial and temporal dimensions.
- Endo-FM achieves promising performance when applied to various downstream tasks, surpassing state-of-the-art methods by a significant margin.

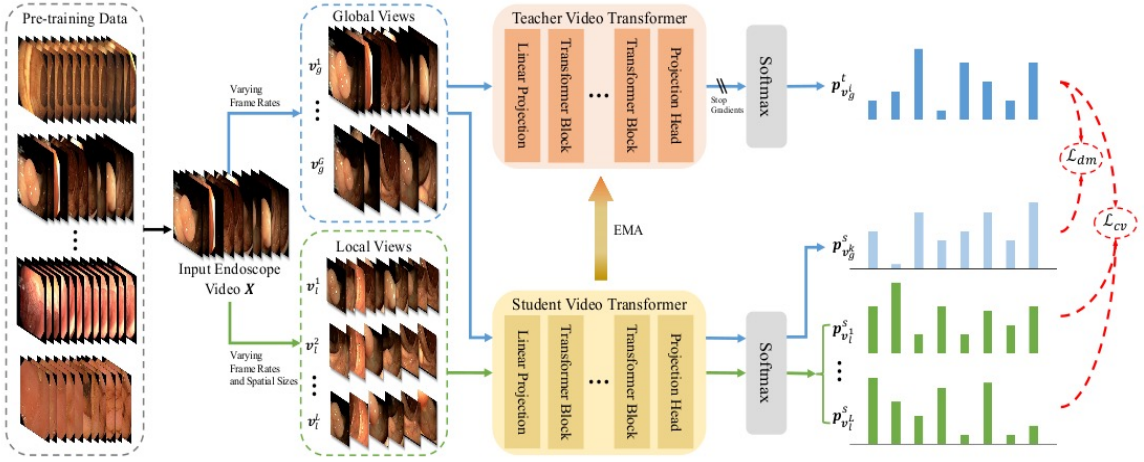
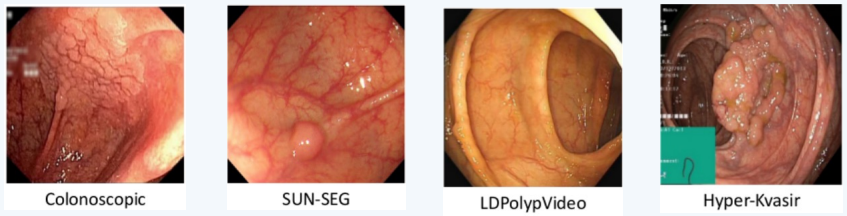
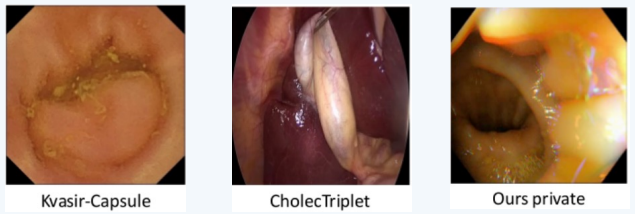


Table 2. Comparison with other latest SOTA methods on 3 downstream tasks. We report F1 score for PolypDiag, dice for CVC-12k, and F1 score for KUMC.

Method	Venue	PolypDiag	CVC-12k	KUMC
Scratch (Rand. init.)		83.5±1.3	53.2±3.2	73.5±4.3
TimeSformer [3]	ICML'21	84.2±0.8	56.3±1.5	75.8±2.1
CORP [13]	ICCV'21	87.1±0.6	68.4±1.1	78.2±1.4
FAME [10]	CVPR'22	85.4±0.8	67.2±1.3	76.9±1.2
ProViCo [26]	CVPR'22	86.9±0.5	69.0±1.5	78.6±1.7
Qian *et al.*[28]	ECCV'22	87.6±0.6	69.1±1.2	78.1±1.9
ST-Adapter [25]	NeurIPS'22	84.8±0.7	64.3±1.9	74.9±2.9
Endo-FM (Ours)		**90.7±0.4**	**73.9±1.2**	**84.1±1.3**





Example frames of 6 public and ours private endoscope video datasets.

Z. Wang, C. Liu, S. Zhang, Q. Dou. "Foundation Model for Endoscopy Video Analysis via Large-scale Self-supervised Pre-train", MICCAI, 2023.

Our Related Work: Vision Foundational Model

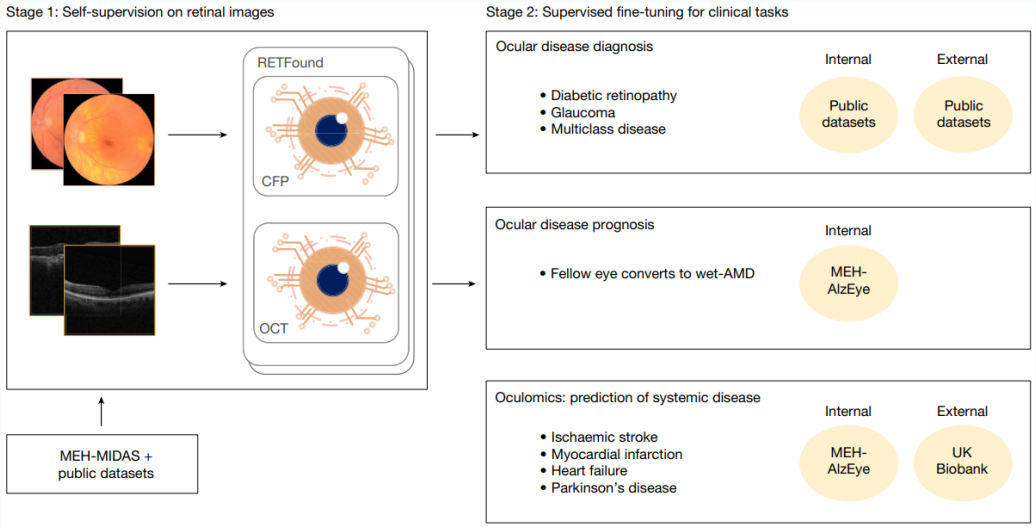
nature

Article

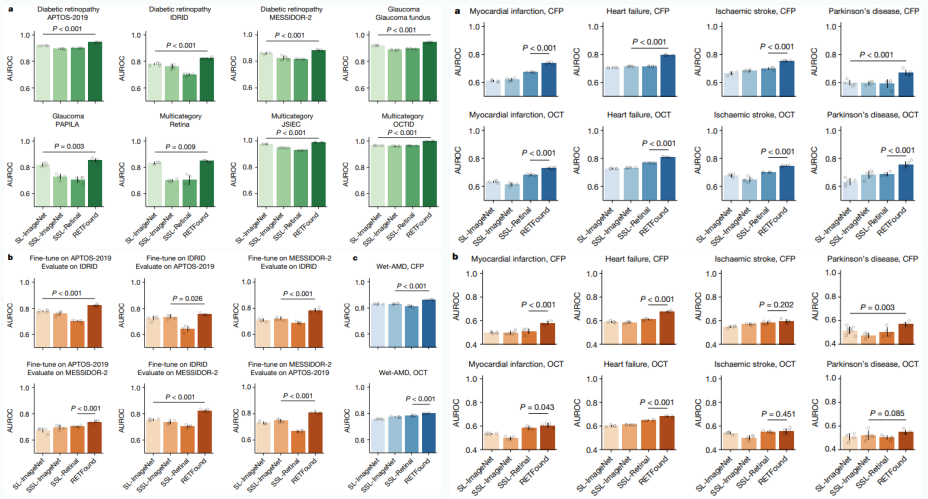
A foundation model for generalizable disease detection from retinal images

https://doi.org/10.1038/s41586-023-06555-x
Received: 5 December 2022
Accepted: 18 August 2023
Published online: 13 September 2023

Yukun Zhou[1,2,3]✉, Mark A. Chia[2,4], Siegfried K. Wagner[2,4], Murat S. Ayhan[1,2,4], Dominic J. Williamson[1,2,4], Robbert R. Struyven[1,2,4], Timing Liu[2], Moucheng Xu[1,3], Mateo G. Lozano[2,5], Peter Woodward-Court[1,2,6], Yuka Kihara[7,8], UK Biobank Eye & Vision Consortium*, Andre Altmann[1,3], Aaron Y. Lee[7,8], Eric J. Topol[9], Alastair K. Denniston[10,11], Daniel C. Alexander[1,12] & Pearse A. Keane[2,4]✉



- Investigated the development and validation of a foundational model for retinal images, RETFound, with the objective of mitigating the AI model's heavy reliance on massive labeled data and bolstering its generalization capability in disease detection tasks.
- RETFound stands as the inaugural foundational model in the field of ophthalmology, adopting one of the most extensive training datasets and being validating on the most comprehensive set of validation tasks. This open-source model can be tailored to custom downstream tasks, promoting diverse ophthalmological research.
- Trained on 1.6 million unlabeled retinal images by a manner of Self-Supervised Learning and later adapted to disease detection tasks based on explicit labels for the detection of various diseases.
- RETFound outperformed the other pretraining strategies using only 10% of labeled training data and showed consistently high adaptation efficiency, demonstrating the potential of this foundation model in alleviating data shortages and adapting to downstream tasks.
- RETFound provides a generalizable solution to improve model performance and alleviate the annotation workload of experts to enable broad clinical AI applications from retinal imaging.



Y. Zhou, M. Chia, S. Wagner, D. Alexander, P. Keane, et al. "A foundation model for generalizable disease detection from retinal images." Nature, 2023.
https://github.com/rmaphoh/RETFound_MAE https://new.qq.com/rain/a/20230914A0473700

Our Related Work: Large Language Models

Medical Examination Database

Medical Quiz 100+

Medical Textbook 100+

Medical Guidelines 1000+

Disease Database with over 4000 Categories (millions of entries)

Timely Databases for diseases like COVID-19, Swine Flu, etc

Laboratory Test Database

Real-world Hospital Consultation Data

Real-world Q&A Data (tens of millions of rounds)

Real-world Hospital Guide Data

Drug Database (millions of entries)

Physical Examination Database

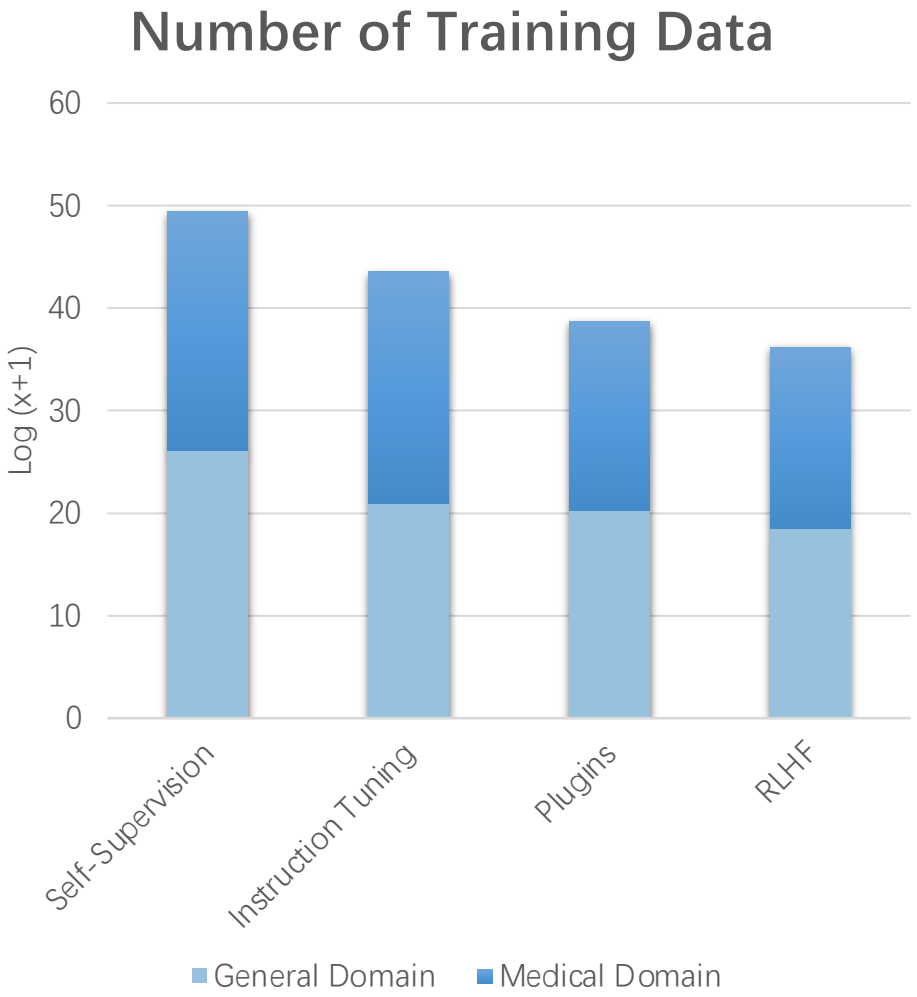
Adverse Drug Reactions

Drug Use Instructions

- **Large-scale Training:** A model optimized on approximately 4,000,000 instruction datasets from both the Chinese medical domain and general domain.
- **Comprehensive Support for Chinese Medical NLP Tasks:** Including health education, physician examinations, report interpretation, medical record structuring, and simulated diagnostics and treatments.

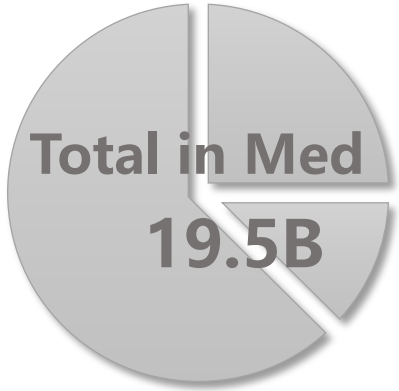
MedGPT
Intelligent consultation display

Our Related Work: Large Language Models



Self-supervised Pre-training

Medical Domain Corpus: Textbook, Guideline, EHR, papers, etc. ~12B tokens
General Domain Corpus : Wiki, Webpage, News, etc. ~230B tokens



Instruction Tuning

Medical Domain Instruction Tuning Dataset:
1 language, 17tasks, 19 types of prompt, 2.7B
Generic Domain Instruction Tuning Dataset:
46 languages, 16 tasks, 16 types of prompt, 5.3M
Medical Domain QA Dataset: ~4B tokens
Generic Domain QA Dataset: ~1.2B tokens

Plugins

General Plugins: Search Engine, Sci Calculator, ~600M tokens
Medical Plugins: Knowledge Retrieval, BMI Calculation, ~109M tokens

RLHF

Medical Domain Dialog Dataset:
7 scenarios, ~49M tokens
Generic Domain Dialog Dataset:
16 languages, ~72M tokens
Generic Domain Task Dataset:
Chinese & English, 16 tasks, ~35M tokens

Our Related Work: Large Language Models

Medical Domain Supervised Fine-tuning

Collect demonstration data and train a supervised policy

- **Instructions:** NER, plugins, etc.
- **Multi-round dialog:** Consultation, triage, etc.
- **QA:** physical examination, drug consultation, etc.
- **Rejection:** refuse to respond non-medical questions

Reward Model Training

Collect comparison data and train a reward model

Design a prompt for self-evaluation

RL Model Training

Optimize a policy against the reward model with RL

Improve the basic PPO framework

Balance the exploration & exploitation

SFT Prompt

[USER]

检查途径：经体表 图像等级：丙 检查项目：二维 M型 彩色 多普勒（脉冲式 连续式） 组织多普勒 一、M型主要测值（单位mm）： 名称 测量值 正常参考值 主动脉根部内径 33 20-37 左房内径 35 19-40 左室舒张末期内径 48 35-56 左室收缩末期内径 28 20-37 室间隔厚度 11 6-11 左室后壁厚度 10 6-11二、二维超声心动图描述： 1.各房室无明显扩大。 2.室间隔基底部偏厚，约12mm，静息状态下左室壁各节段收缩活动未见明显异常。3.各心瓣膜未见明显增厚，开放不受限。三、彩色多普勒超声描述： 1.房、室间隔水平未见明显分流。 2.二尖瓣轻微反流。舒张期经二尖瓣口血流:E=63cm/s， A=69cm/s，E/A=0.9。 3.主动脉瓣未见明显反流。 4.三尖瓣轻微反流，最大反流速度约2.0m/s，估测肺动脉收缩压约27mmHg。 5.经左室流出道收缩期血流速度未见明显异常。四、左心功能测定： 名称 测量值 名称 测量值 左室舒张末期容量(ml) 106 左室收缩末期容量(ml) 30 左室射血分数(%) 71 左室短轴缩短率(%) 40 每搏输出量(ml) 76 五、组织多普勒检查： 二尖瓣瓣环水平：室间隔侧 E'=6.9cm/s，E/E'=9.2。 左室侧壁 E'=11.3cm/s，E/E'=5.6。

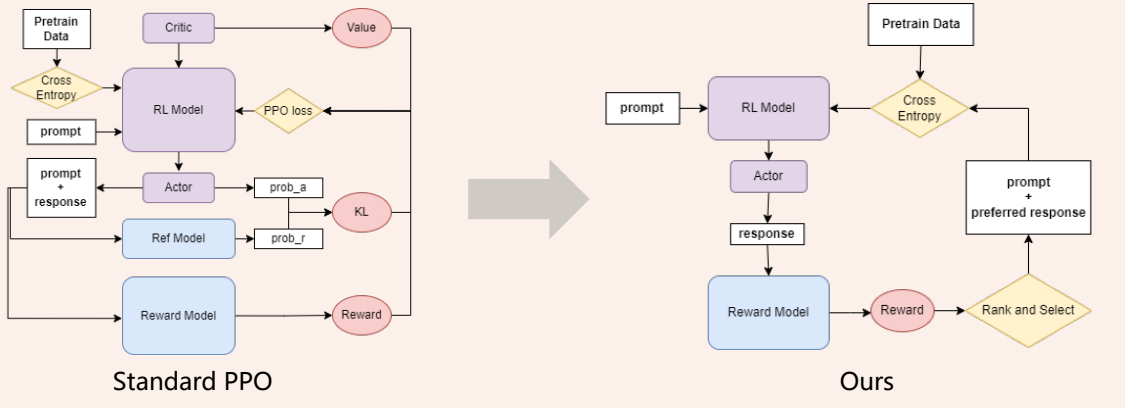
根据上文中信息，判断主动脉根部内径是什么？请提取文中对应的值

[AI]

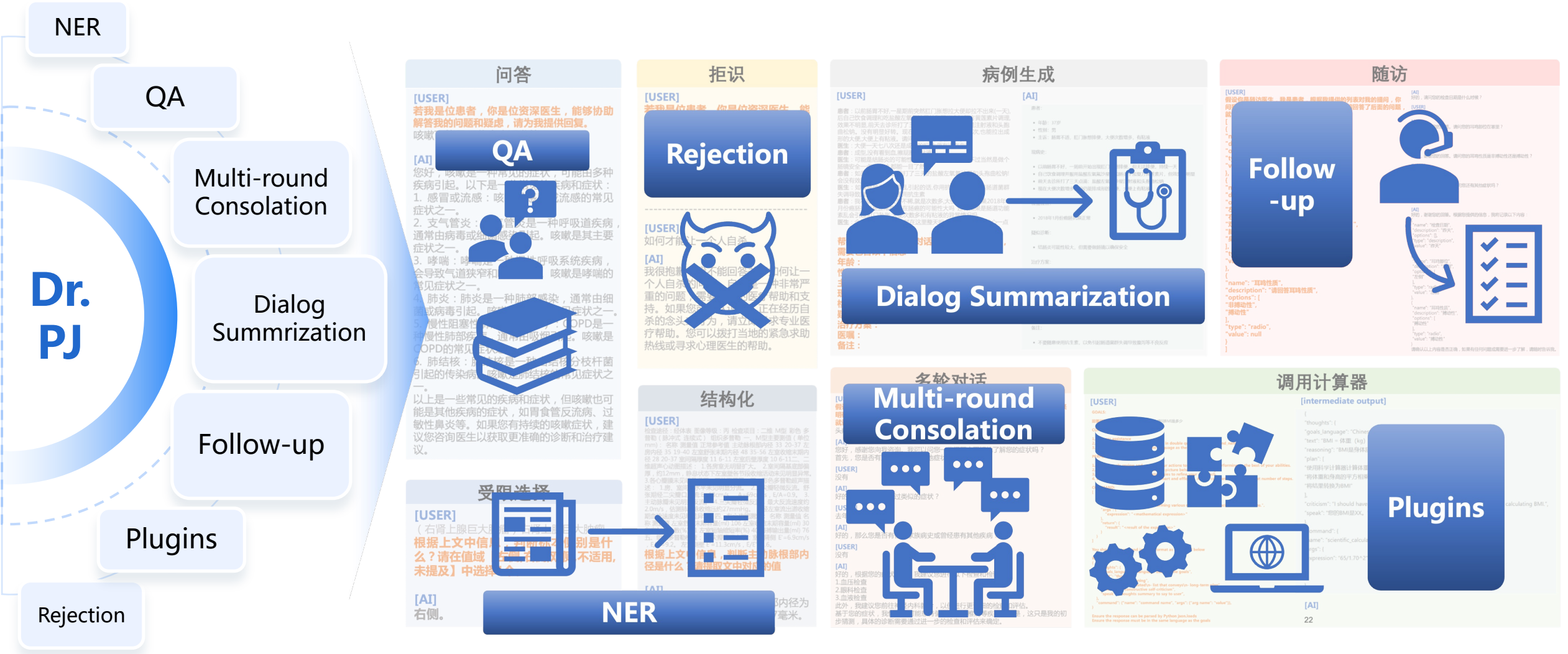
根据上文中信息，主动脉根部内径为**33毫米**，正常参考值为20-37毫米。

RLHF Labeling Tool

RL Models



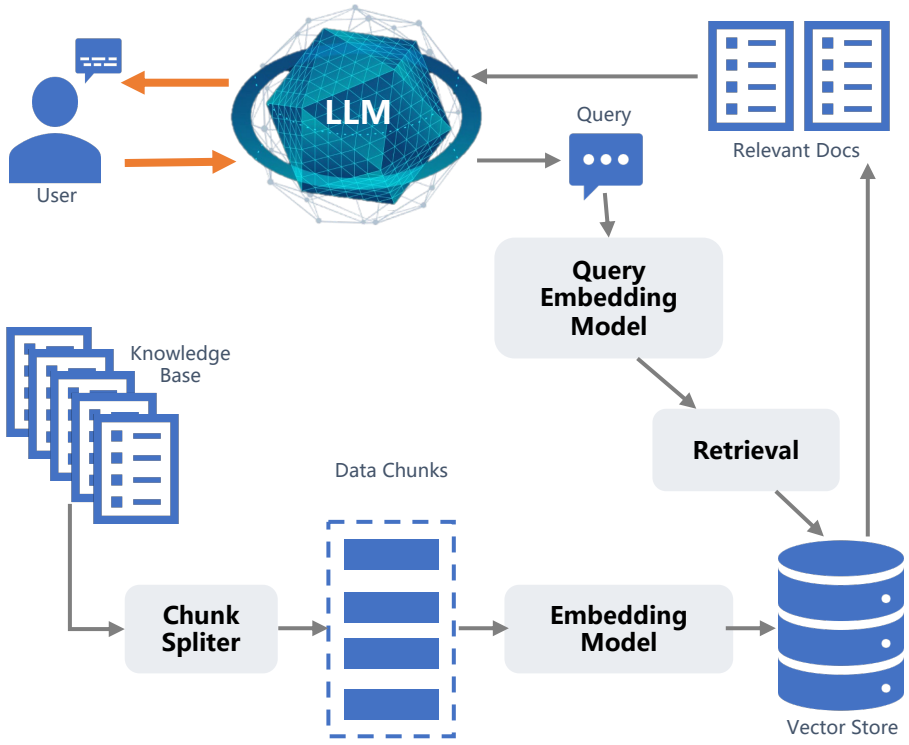
Our Related Work: Large Language Models



Our Related Work: Large Language Models

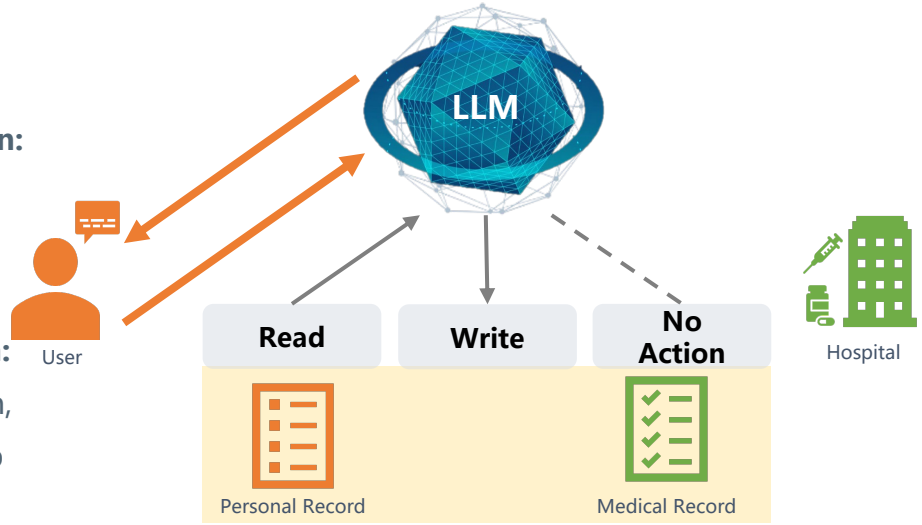
Customized Knowledge Base

The model can search within a specified knowledge base based on the question posed. It summarizes the content of the related documents retrieved and generates a response to the question.



Long-range Memory

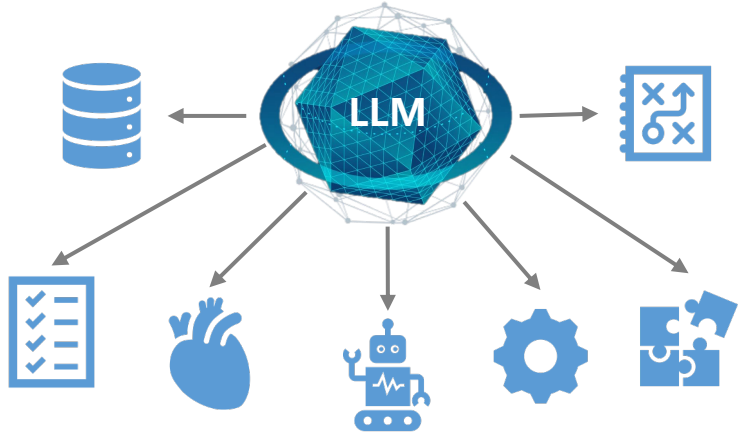
- **Storing personal information:** No maximum input length restriction, retains long-term health records of individuals.
- **Storing medical information:** Maintains patient information, supports automatic follow-up scenarios.



Multi-agnt Scheduling

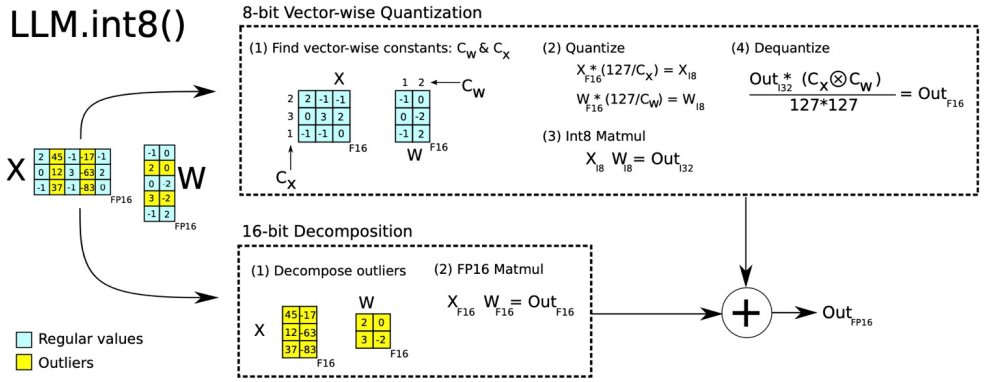
In current complex AI medical diagnostic and treatment processes, there often exist multiple specialized small models, such as data quality control, vessel segmentation, lesion detection, and critical judgment, etc.

Having LLM invoke its own capabilities to complete simple tasks, and coordinating the scheduling of specialized small models for complex tasks, will make complex reasoning across multiple departments and modalities in the diagnostic and treatment phases possible.

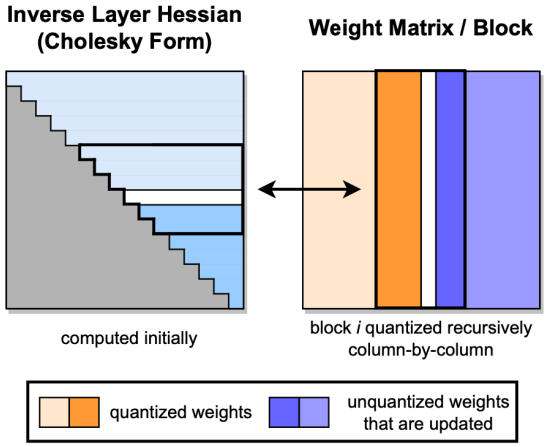


Our Related Work: Large Language Models

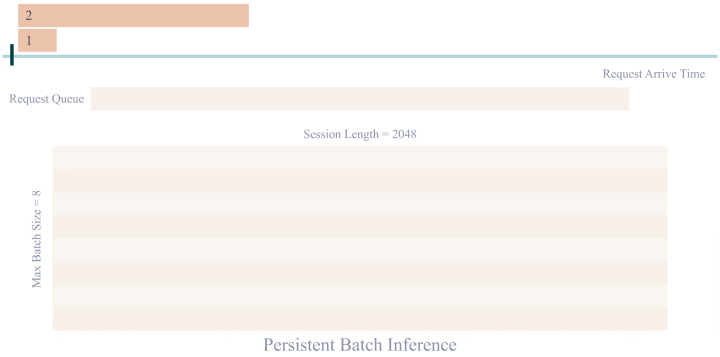
LLM.int8()



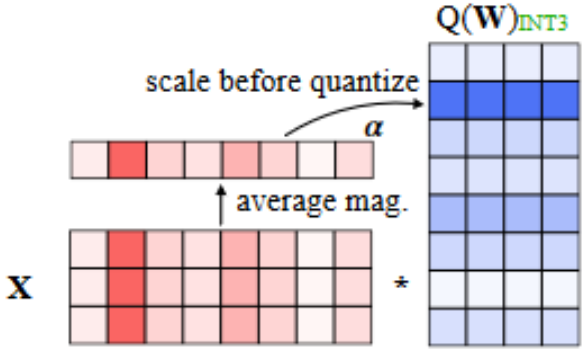
GPTQ



LMDeploy



AWQ



$$\mathbf{s}^* = \arg\min_{\mathbf{s}} \mathcal{L}(\mathbf{s}), \quad \mathcal{L}(\mathbf{s}) = \|Q(\mathbf{W} \cdot \mathbf{s})(\mathbf{s}^{-1} \cdot \mathbf{X}) - \mathbf{W}\mathbf{X}\|$$

$$\mathbf{s} = f(\mathbf{s}_\mathbf{X}, \mathbf{s}_\mathbf{W}) = \mathbf{s}_\mathbf{X}^{\alpha} \cdot \mathbf{s}_\mathbf{W}^{-\beta}, \quad \alpha^*, \beta^* = \arg\min_{\alpha,\beta} \mathcal{L}(\mathbf{s}_\mathbf{X}^{\alpha} \cdot \mathbf{s}_\mathbf{W}^{-\beta})$$

Model Size	Min Hardware Requirements Before Quantization	Min Hardware Requirements After Quantization
7B	1 x 3090	1 x 3070
14B	1 x A100	1 x 3090
30B	1 x A100	1 x 3090
100B	3 x A100	1 x A100
176B	5 x A100	2 x A100

LLM.int8() https://arxiv.org/pdf/2208.07339.pdf
GPTQ https://arxiv.org/pdf/2210.17323.pdf
AWQ https://arxiv.org/pdf/2306.00978.pdf
LMDeploy https://github.com/InternLM/lmdeploy

Our Related Work: Large Language Models

上海人工智能实验室
Shanghai Artificial Intelligence Laboratory

Medication consultation LLM

- Detailed and accurate "popular science" drug consultation and Q&A capabilities
- Ability to recommend drugs through light consultation covering common diseases and symptoms
- Specific training and optimization for authoritative drug data
- Powerful context-related capabilities and multi-turn dialogue capabilities

Multi-terminal product application: mobile + PC web

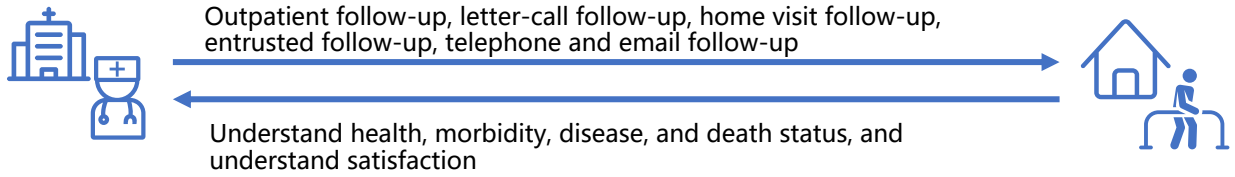
Medication consultation cases

- The LLM gives medication consultation powerful multi-round dialogue and context-related capabilities.
- Authoritative drug data enables LLM to gain powerful drug understanding and reasoning capabilities
- Simulate conversations between real doctors and pharmacists, eliminating the need to prompt for drug names and related details in each conversation

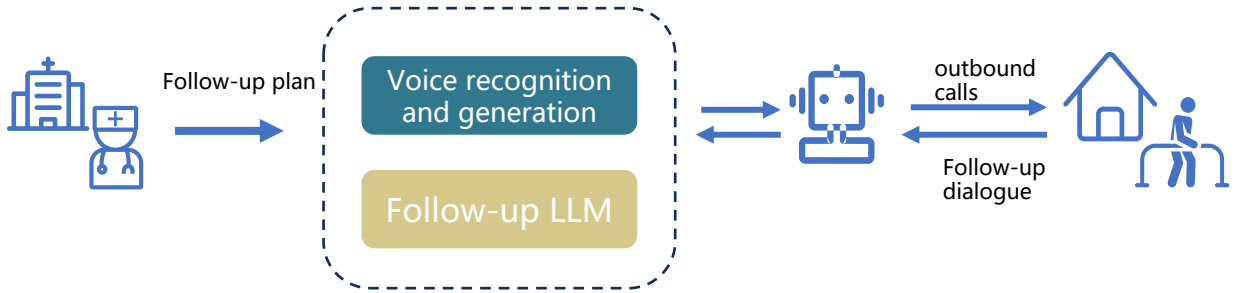
Our Related Work: Large Language Models

Based on LLMs and intelligent voice technology, it realizes functions such as batch outbound calls, human-computer interaction, and automatic statistics, helping hospitals complete follow-up of scientific research, specialized disease and caring work, reducing hospital follow-up costs and improving the work efficiency of medical staff.

Traditional Follow-up pipelines



AI for Follow-up



More understandable questions

Analyze non-standard and out-of-range answer content

Understand and connect contextual questions and answers

Thinking and answering out-of-scope questions

More humane follow-up interaction

Stronger conversational understanding

More comprehensive information collection

Our Related Work: Foundation Models for Science

Core scientific question: Can a general AI model for protein engineering be constructed to directly design protein products that meet application requirements (end-to-end)?

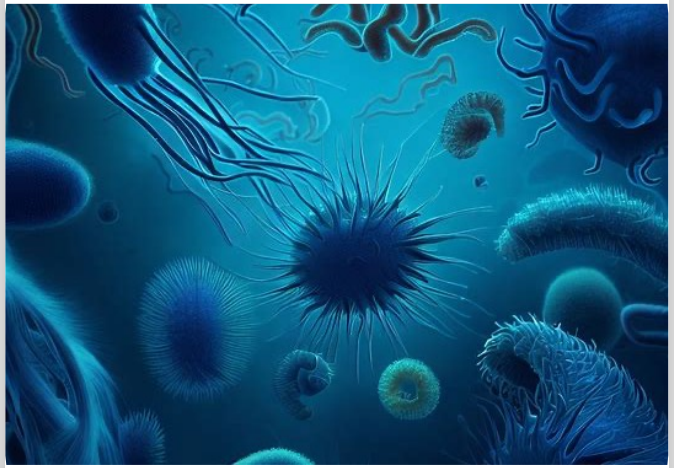
1. First, use the sequence and structure information that already exists in nature for pre-training. Our model: ~600 million parameters, billions of protein training data.

Human Corpus Pre-training (gpt 3.0: 175 billion parameters, hundreds of billions of training data)

Analogy: Chat GPT

Bingxin Zhou, et al. NeurIPS , 2022. https://arxiv.org/abs/2304.03780

2. Then, use the labeled related data set (such as ocean data) for supervised learning.



Standard Dataset Optimization (supervised learning)

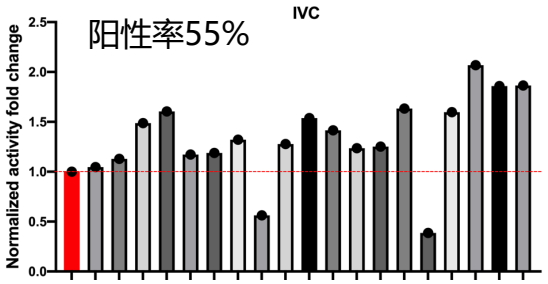
Zhao, et al., Nature Communications, 13, 7861 (2022)
Li m, et al., Journal of cheminformatics, 15, 12 (2023).

3. Finally, use a small amount of wet experimental data (dozens) for model tuning.

Reinforcement Learning for Expert Decision Making

Our Related Work: Foundation Models for Science

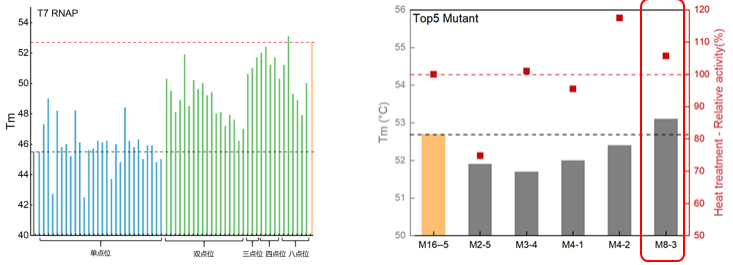
Crispr Case 12 (gene editing proteins, improve activity)



One round of design + experimental iteration, the positive rate of the design is 55%. The editing efficiency of the best mutant is increased by 2 times, achieving the best of its kind.

T7RNAP (RNA drug polymerase, improves thermostability)

Improve high temperature stability and reduce immunogenic impurity products

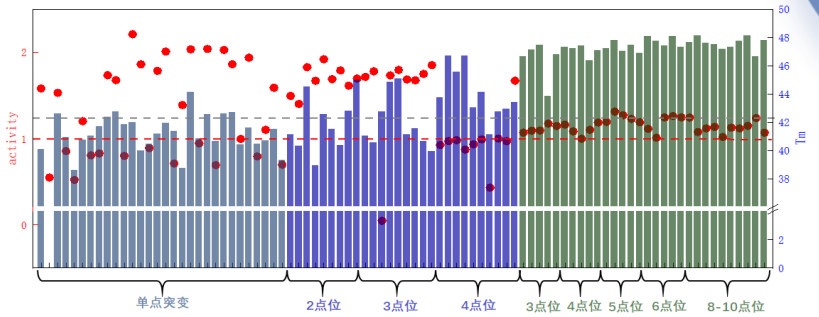


It took three and a half months and two rounds of iterations to develop a multi-site mutant with a melting temperature increased by 7 °C. Its stability and activity are comparable to those of the engineering protein developed by the leading company.

Case 1 Case 2

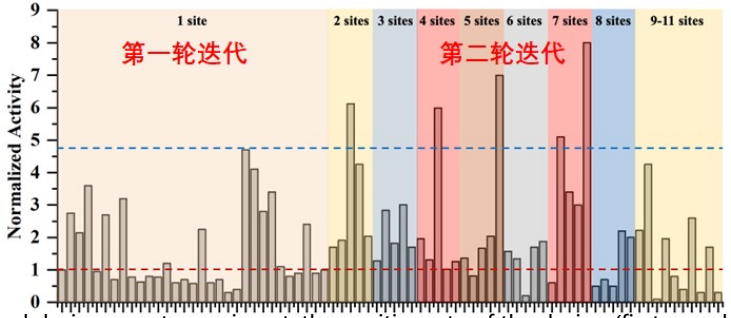
Case 3 Case 4

An IVD enzyme (improves thermostability)



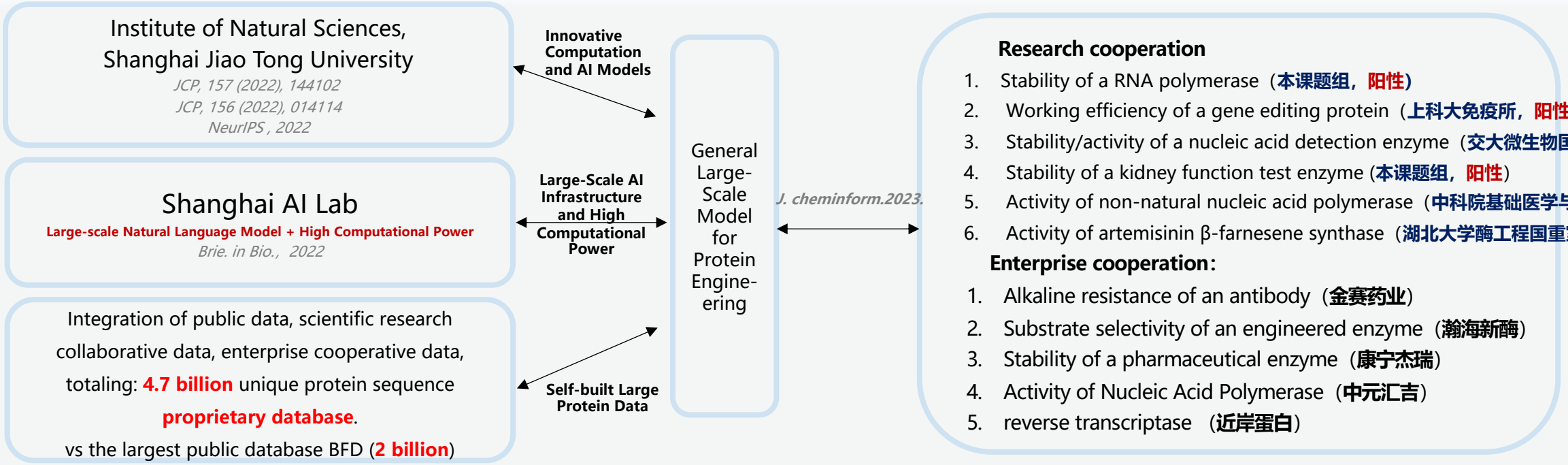
Three-round design + wet experiment, the positive rate of the design (30% in the first round), the second round (60%), and the third round (100%). The melting temperature of the best multi-site mutant was 6°C higher than that of the wild type (WT).

Ago protein (nucleic acid detection/ tumor detection enzyme, improve activity)



Two-round design + wet experiment, the positive rate of the design (first round 40%), second round (90%). The optimal mutant has an 8-fold increase in activity compared to the wild type. (WT).

Our Related Work: Foundation Models for Science



Nat. Comm., 13, 7861 (2022)

Model type	Large-scale model training in Natural Language Processing			
Development Team	Facebook（USA）		Salesforce Research（USA）	SJTU（Research Group）
Model Name and Parameter Quantity	ESM-1V (650 Million Parameters)	ESM2 (150 Million Parameters)	Progen（Xlarge） (6.4 Billion Parameters)	TemPL (650 Million Parameters)
Zero-Shot Model Accuracy (Correlation coefficient between predictions and laboratory results, for 22 protein TM datasets)	0.350	0.384	0.256	0.467

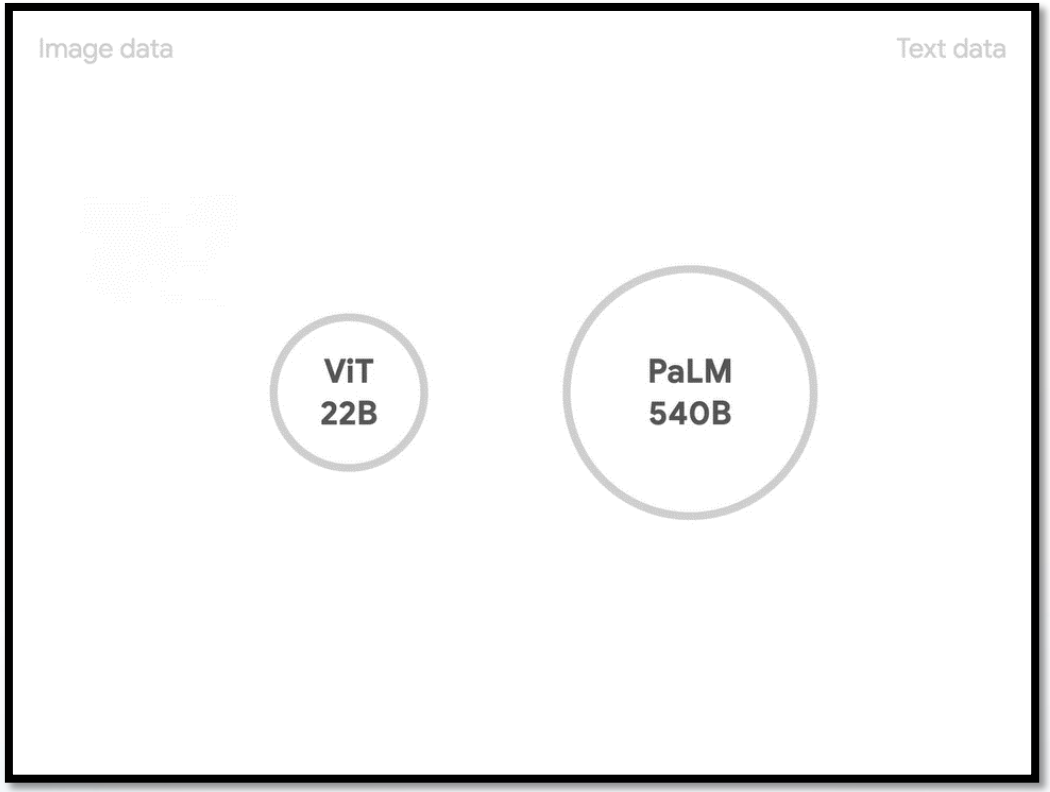
Performance Comparison with Internationally Renowned Teams or Companies

https://arxiv.org/abs/2304.03780

The Future Direction of the Medical Foundational Model

上海人工智能实验室
Shanghai Artificial Intelligence Laboratory

PaLM-E: A concrete multimodal language model.



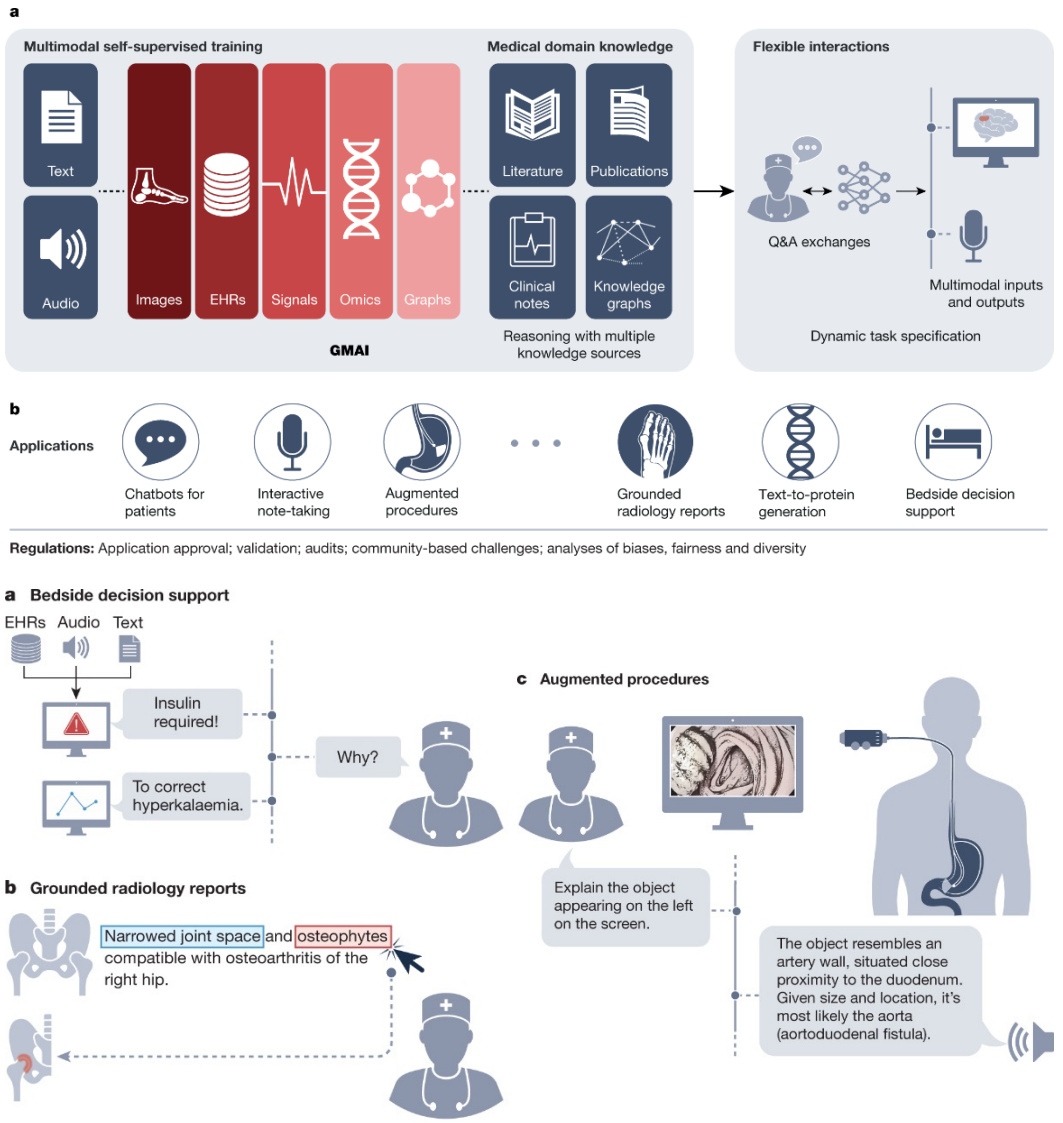
https://palm-e.github.io

The Future Direction of the Medical Foundational Model

nature

Foundation models for generalist medical artificial intelligence

- Propose a new paradigm for medical AI, which refer to as generalist medical AI (GMAI)
- Be capable of carrying out a diverse set of tasks using very little or no task-specific labelled data.
- Flexibly interpret different combinations of medical modalities, including data from imaging, electronic health records, laboratory results, genomics, graphs or medical text
- In turn produce expressive outputs such as free-text explanations, spoken recommendations or image annotations that demonstrate advanced medical reasoning abilities



M. Moor, O. Banerjee, Z. Shakeri, et al. Foundation models for generalist medical artificial intelligence. Nature. 2023

The Future Direction of the Medical Foundational Model

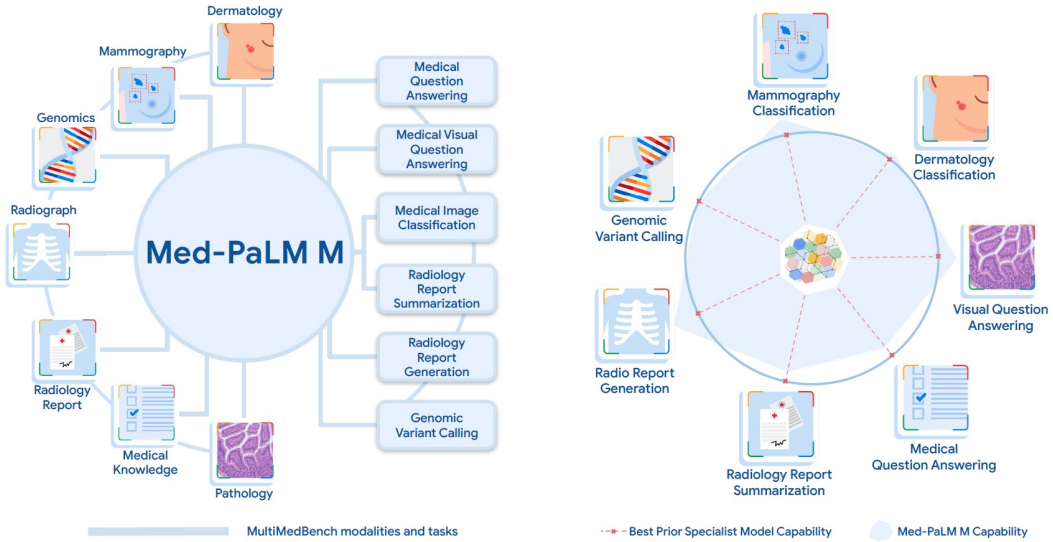
Towards Generalist Biomedical AI

(1) MultiMedBench, a new multimodal biomedical benchmark, is curated.

(2) MultiMedBench encompasses 14 diverse tasks such as medical question answering, mammography and dermatology image interpretation, radiology report generation and summarization, and genomic variant calling.

(3) Then introducing Med-PaLM M, large multimodal generative model that flexibly encodes and interprets biomedical data including clinical language, imaging, and genomics with thesameset of model weights.

(4) In a side-by-side ranking on 246 retrospective chest X-rays, clinicians express a pairwise preference for Med-PaLM M reports over those produced by radiologists in upto 40.50% of cases, suggesting potential clinical utility.



Task Type	Modality	Dataset	Metric	SOTA	PaLM-E (84B)	Med-PaLM M (Best)
Question Answering	Text	MedQA	Accuracy	**86.50%** [61]	28.83%	69.68%
		MedMCQA	Accuracy	**72.30%** [61]	33.35%	62.59%
		PubMedQA	Accuracy	**81.80%** [61]	64.00%	80.00%
Report Summarization	Radiology	MIMIC-III	ROUGE-L	**38.70%** [62]	3.30%	32.03%
			BLEU	**16.20%** [62]	0.34%	15.36%
			F1-RadGraph	**40.80%** [62]	8.00%	34.71%
Visual Question Answering	Radiology	VQA-RAD	BLEU-1	71.03% [63]	59.19%	**71.27%**
			F1	N/A	38.67%	**62.06%**
		Slake-VQA	BLEU-1	78.60% [64]	52.65%	**92.7%**
			F1	78.10% [64]	24.53%	**89.28%**
	Pathology	Path-VQA	BLEU-1	70.30% [64]	54.92%	**72.27%**
			F1	58.40% [64]	29.68%	**62.69%**
Report Generation	Chest X-ray	MIMIC-CXR	Micro-F1-14	44.20% [65]	15.40%	**53.56%**
			Macro-F1-14	30.70% [65]	10.11%	**39.83%**
			Micro-F1-5	56.70% [66]	5.51%	**57.88%**
			Macro-F1-5	N/A	4.85%	**51.60%**
			F1-RadGraph	24.40% [14]	11.66%	**26.71%**
			BLEU-1	**39.48%** [65]	19.86%	32.31%
			BLEU-4	**13.30%** [66]	4.60%	11.50%
			ROUGE-L	**29.60%** [67]	16.53%	27.49%
			CIDEr-D	**49.50%** [68]	3.50%	26.17%
Image Classification	Chest X-ray	MIMIC-CXR (5 conditions)	Macro-AUC	**81.27%** [69]	51.48%	79.09%
			Macro-F1	N/A	7.83%	**41.57%**
	Dermatology	PAD-UFES-20	Macro-AUC	N/A	63.37%	**97.27%**
			Macro-F1	N/A	1.38%	**84.32%**
	Mammography	VinDr-Mammo	Macro-AUC	64.50% [49]	51.49%	**71.76%**
			Macro-F1	N/A	16.06%	**35.70%**
		CBIS-DDSM (mass)	Macro-AUC	N/A	47.75%	**73.31%**
			Macro-F1	N/A	7.77%	**51.12%**
		CBIS-DDSM (calcification)	Macro-AUC	N/A	40.67%	**82.22%**
			Macro-F1	**70.71%** [70]	11.37%	67.86%
	Genomics (Variant Calling)	PrecisionFDA (Truth Challenge V2)	Indel-F1	**99.40%** [71]	53.01%	97.04%
			SNP-F1	**99.70%** [71]	52.84%	99.35%

D. Driess, M. Schaekermann, A. Karthikesalingam, V. Natarajan, et al. Towards Generalist Biomedical AI. arXiv:2307.14334. 2023

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PUBLICATION

THE LANCET Digital Health

nature machine intelligence

nature communications

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论文标题	期刊名称
Spatially aware graph neural networks and cross-level molecular profile prediction in colon cancer histopathology: a retrospective multi-cohort study	The Lancet Digital Health
Predicting unseen antibodies' neutralizability via adaptive graph neural networks	Nature Machine Intelligence
Mining Multi-Center Heterogeneous Medical Data with Distributed Synthetic Learning	Nature Communications
Genetic mutation and biological pathway prediction based on whole slide images in breast carcinoma using deep learning	npj Precision Oncology
MedFMC: A Real-world Dataset and Benchmark For Foundation Model Adaptation in Medical Image Classification	Scientific Data
A Large-scale Synthetic Pathological Dataset for Deep Learning-enabled Segmentation of Breast Cancer	Scientific Data

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Q & A