

# CSE 428 Computational Biology Capstone

# What is research

- Engineering: Solve an existing problem
- Research: Propose a new problem
  - ✓ Define the problem
    - Problem setting (input, output)
    - Benchmark for validation
    - Baselines
  - ✓ Propose the first solution (which might not be a good solution)
  - ✓ This problem cannot be trivial (Significance)
  - ✓ This problem has not been studied before (Novel)
    - ✓ Why people have never studied it before?
  - ✓ Solving this problem is feasible (Feasible)

# Why interdisciplinary research is impactful

- Lots of opportunities
- New methods from domain A can solve important but unaddressed problems in domain B.
  - Apply sequence modeling to study protein sequence (protein sequence is similar to text sequence)
  - Train foundation model/LLM to study protein sequence
  - Train multi-modal LLM to jointly study protein structure and protein sequence.

# Borrow ideas from other domains

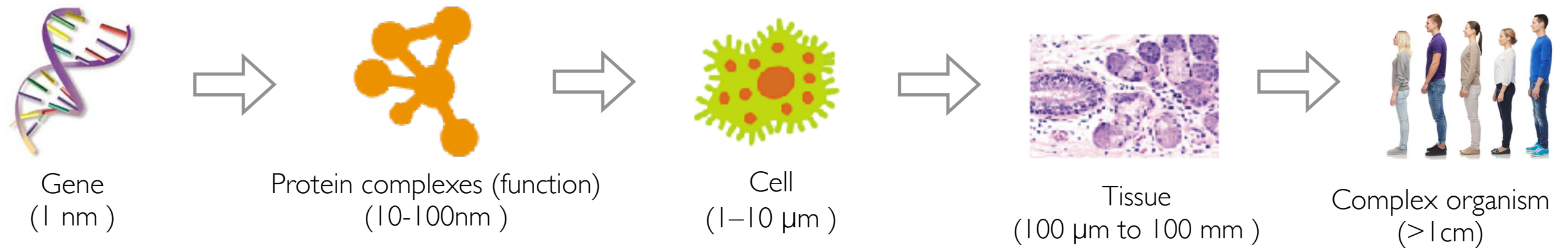
- From less novel to very novel
  - Use a method in one comp bio problem (e.g., protein structure) to solve another comp bio problem (e.g., chromatin structure)
  - Use a new ML technique from CV/NLP to solve a comp bio problem (CSE428 will focus on this)
  - Use a method in other subjects (quantum computing, theoretical physics) to solve a comp bio problem



# What can you learn from research

- Career path for a CS student
  - SDE (no need to have PhD degree):
    - Prompt GPT4 for something.
  - Machine learning engineer (PhD degree is a big plus)
    - Fine-tune GPT4 for something: where to get the fine-tune data, which fine-tune techniques to use
    - Need substantial research ability to understand and modify existing algorithm
  - Research scientist/faculty (PhD is required)
    - Develop OpenAI GPT4
    - It is all about research
- CSE 428 will be at the scope of Machine learning engineer

# CSE427: Computational methods for biology at different scales



A rich hierarchy of biological subsystems at multiple scales: genotypic variations in nucleotides (1 nm scale)  $\rightarrow$  proteins (1-10 nm)  $\rightarrow$  protein complexes (10-100 nm), cellular processes (100 nm)  $\rightarrow$  phenotypic behaviors of cells (1-10  $\mu\text{m}$ ), tissues (100  $\mu\text{m}$  to 100 mm),  $\rightarrow$  complex organisms (>1 m).

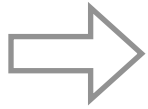
source: Yu, Michael Ku, et al. "Translation of genotype to phenotype by a hierarchy of cell subsystems." *Cell systems* 2.2 (2016): 77-88.

# Computational methods for biology at different scales



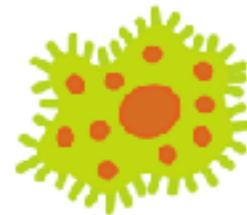
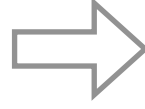
Gene  
(1 nm)

Genetics



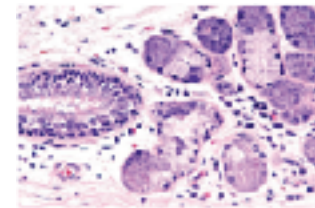
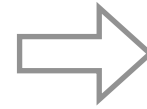
Protein complexes (function)  
(10-100nm)

Systems biology



Cell  
(1-10  $\mu$ m)

Cellular biology



Tissue  
(100  $\mu$ m to 100 mm)

Medical imaging



Complex organism  
(> 1cm)

Computational medicine

Focus of CSE 427

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# CSE 428 projects

- Option 1: define your own project and work by yourself
- Option 2: work on a suggested topic
  - Option 2A: pick a paper from Su-In Lee, Sara Mostafavi, Bill Noble
  - Option 2B: pick a paper from my lab (will introduce later)
  - Understand everything of the paper!
  - Reproduce some key results
  - Try to improve and modify the model

# What you need to submit

- A course report
- A Jupyter notebook commit to the course GitHub
  - Mark down of a pipeline from reading data to produce prediction/analyses.

# Schedule and grading

1. Three presentations
2. course project presentation (60%), final report (40%)

## Tentative Schedule

Date	Topic
4/1	Welcome/overview. Introduction to CSE428. (Sheng)
4/8	Project topic presentation (first half)
4/15	Project topic presentation (second half)
4/22	Working time
4/29	Mid-term project idea presentation (first half)
5/6	Mid-term project idea presentation (second half)
5/13	Working time
5/20	Working time
5/27	Final project presentation (first half)
6/3	Final project presentation (second half)

# First presentation: decide on a paper

- Choose one paper
- Focus on the significance
  - Why is this problem important
  - Why are you interested in this project.
- 5 minutes per student.
- More instruction later based on the number of students



# How to read papers

- This is the most important thing
  - One researcher spends 3 hours per day to read paper
- Read new papers (2023-) and papers published in top journals/conferences
  - Nature, Science, Cell, Recomb, ISMB
- Tips for how to read papers
  - Focus less on the background, introduction and motivation
  - Focus more on the problem setting (input, output)
  - Focus more on the datasets for evaluation (size, accessible, simulated data or real data)
- The more you read, the faster you will read.
- Ultimate goal: you can “predict” what method/dataset/evaluation this paper will do after reading the abstract

# Pick the problem to work on

- Only work on the frontier methods
- Don't be ambitious!
- Only work on problem that an existing paper has worked on and you can fully understand that paper
- Only work on problem that you can clearly evaluate

# Pick the problem to work on

- Pick a paper from Profs from Allen School
  - Su-In Lee, Sara Mostafavi, Sheng Wang, Bill Noble
- Read at least three papers in the first week
  - Each student will do a presentation next week
  - Significance
  - Innovation
  - Methods and evaluation
  - Why do you like this paper?

# How to borrow ideas from other domains

- Find the commonality by understanding the problem setting
  - Problem A in computer vision has the same problem setting as Problem B in comp bio.
  - Talk to people work in CV/NLP

# Experiments

1. Don't implement your ideas first
2. First, reproduce the results from baselines
3. Test baselines on new datasets and examine the performance
4. If the performance is bad, figuring out the reason

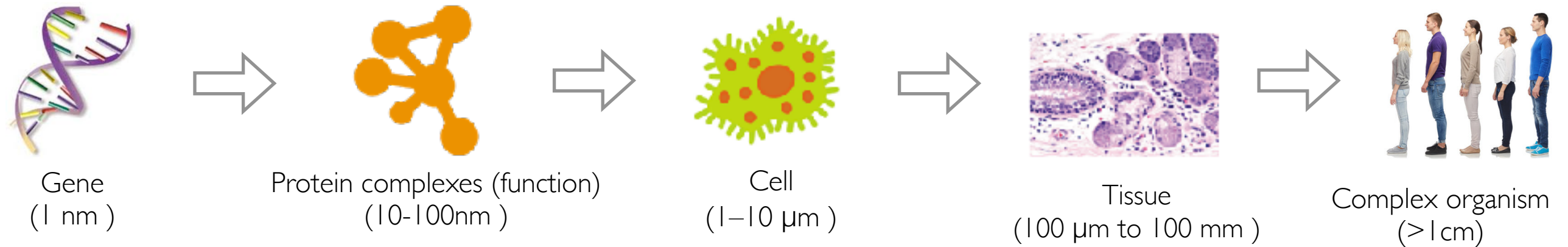
# Uniqueness of comp bio research

- Interdisciplinary subject
  - Communication is very important
  - Understand biologists/doctors' needs
  - Clearly explain our methods
- New subject
  - Lots of opportunities
  - Less well-defined benchmarks

# Reflection on my research career

- 2009-2013 Undergrad in CS, Peking university
- 2013-2018 PhD in CS, UIUC
- 2018-2020 Postdoc in Medicine, Stanford
- Most important thing is choosing the research area.

# CSE427: Computational methods for biology at different scales



A rich hierarchy of biological subsystems at multiple scales: genotypic variations in nucleotides (1 nm scale)  $\rightarrow$  proteins (1-10 nm)  $\rightarrow$  protein complexes (10-100 nm), cellular processes (100 nm)  $\rightarrow$  phenotypic behaviors of cells (1-10  $\mu\text{m}$ ), tissues (100  $\mu\text{m}$  to 100 mm),  $\rightarrow$  complex organisms (>1 m).

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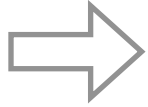


# Computational methods for biology at different scales



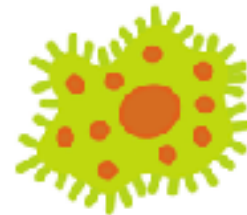
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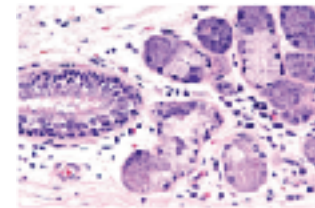
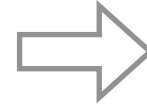
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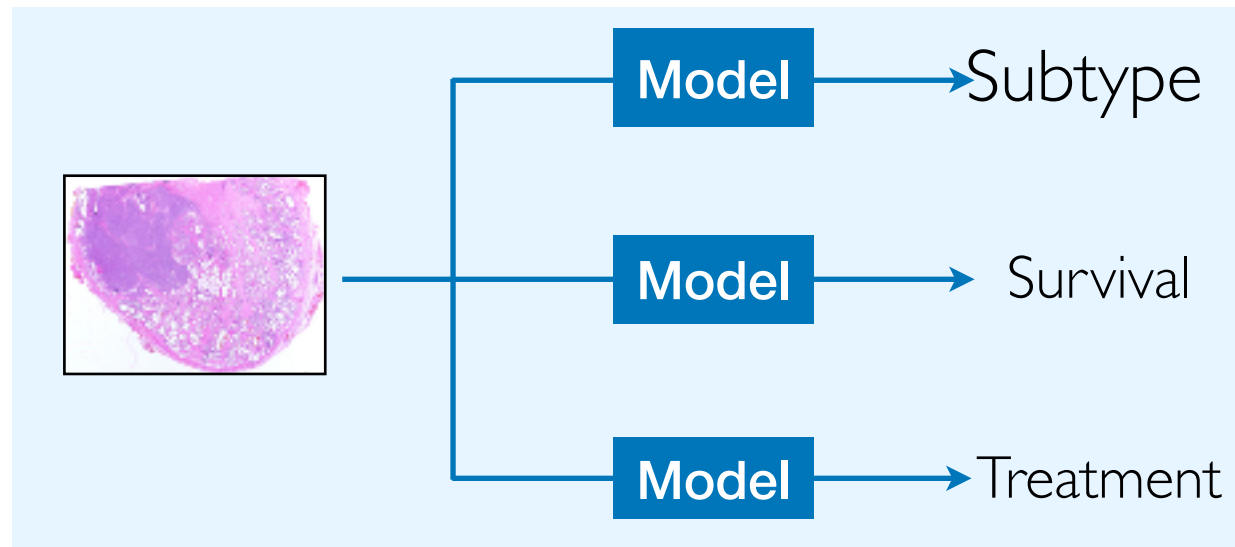
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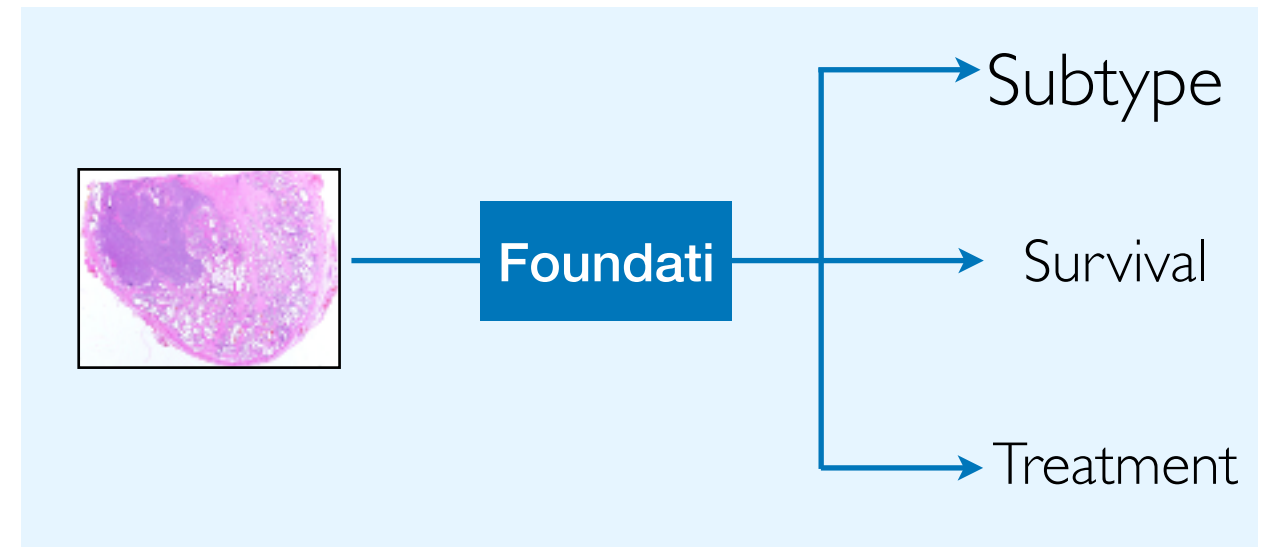
Focus of CSE 427

# Four paradigms in AI for Medicine

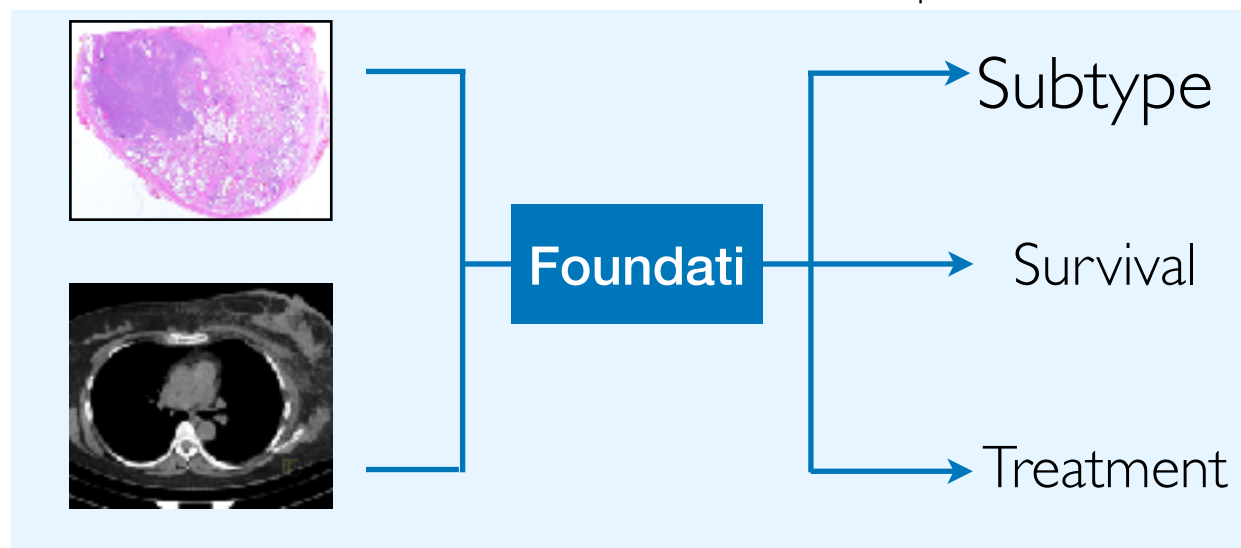
Deep learning (2012)  
One model for one task



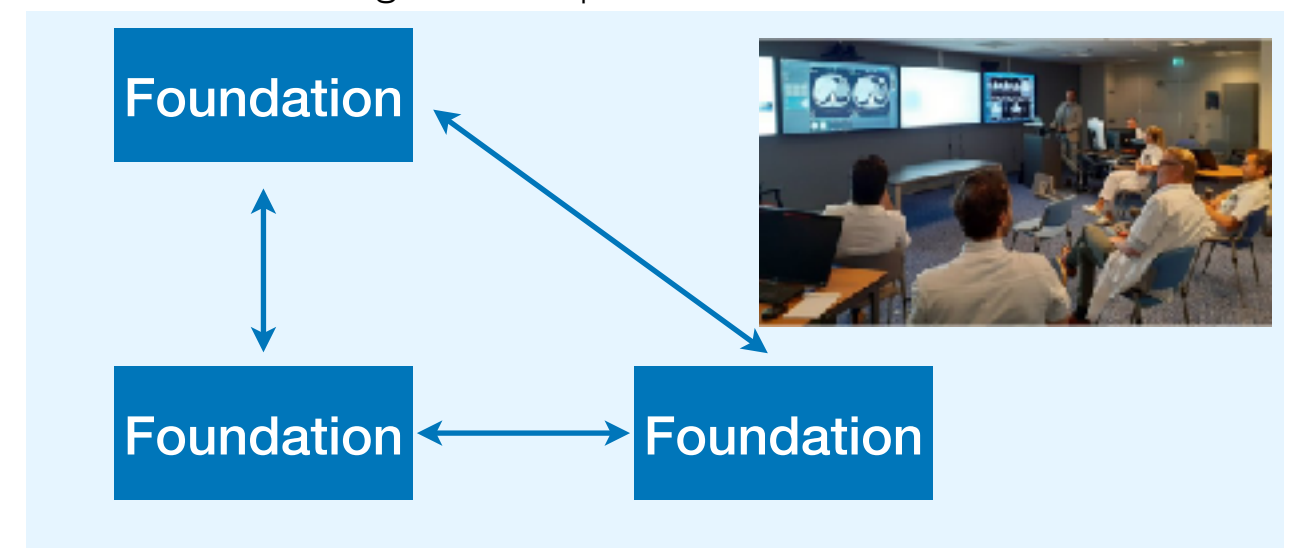
Foundation model (2022)  
One model for all tasks



Multi-modal foundation model (2023)  
One model takes different inputs

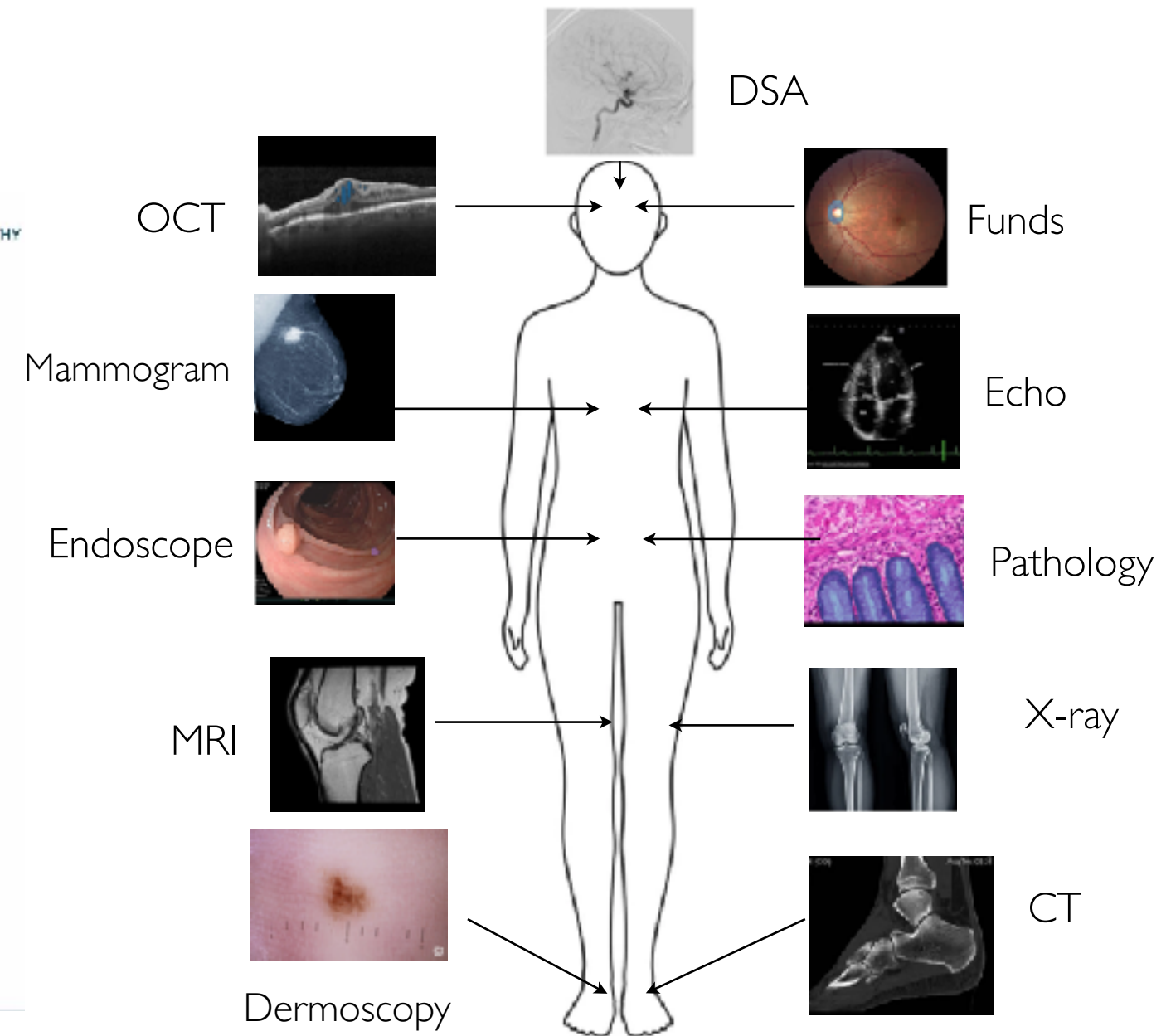
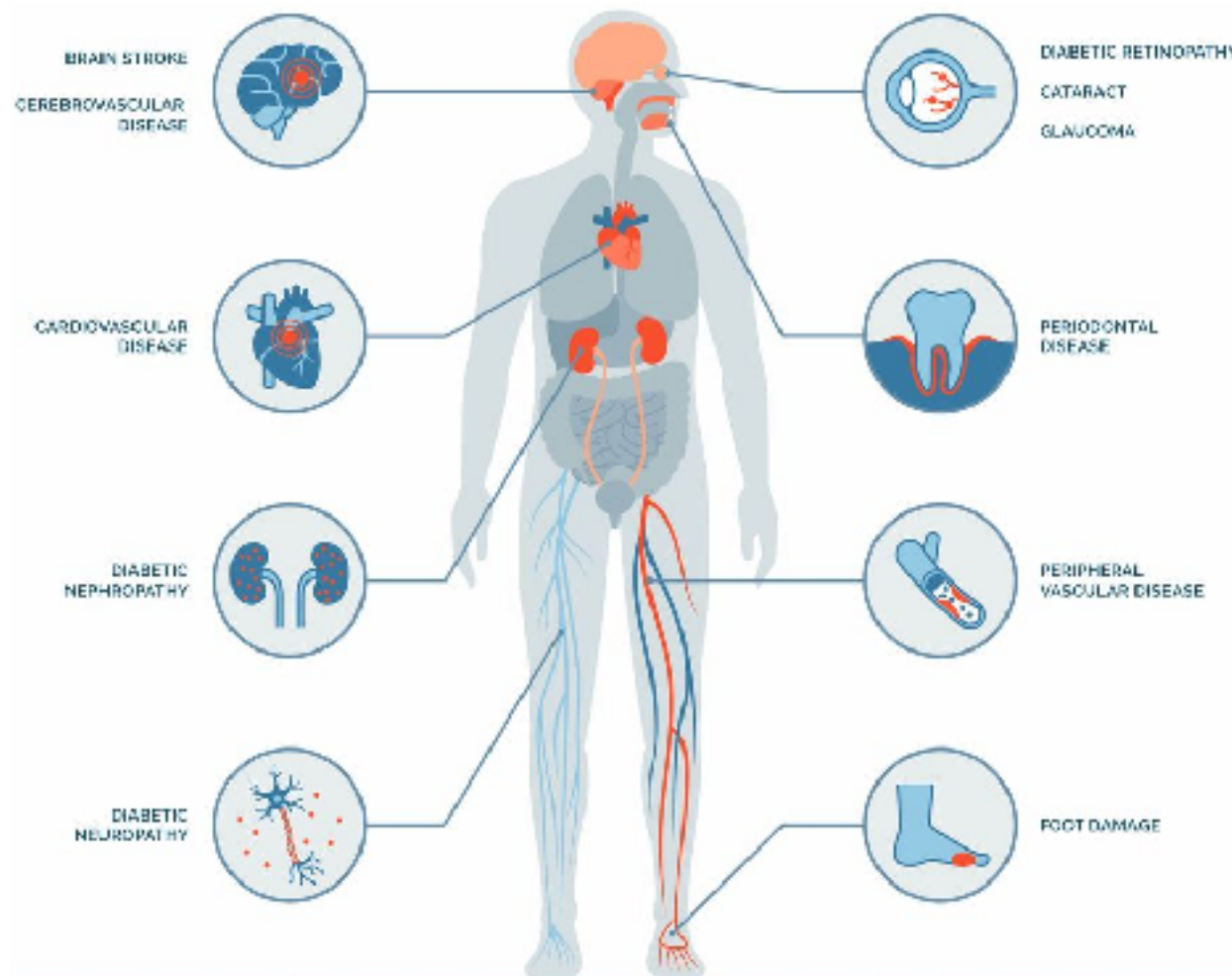


Multi-agent (2024)  
Integrate multiple foundation models



# Medicine is inherently multi-modal

## Complication of diabetes

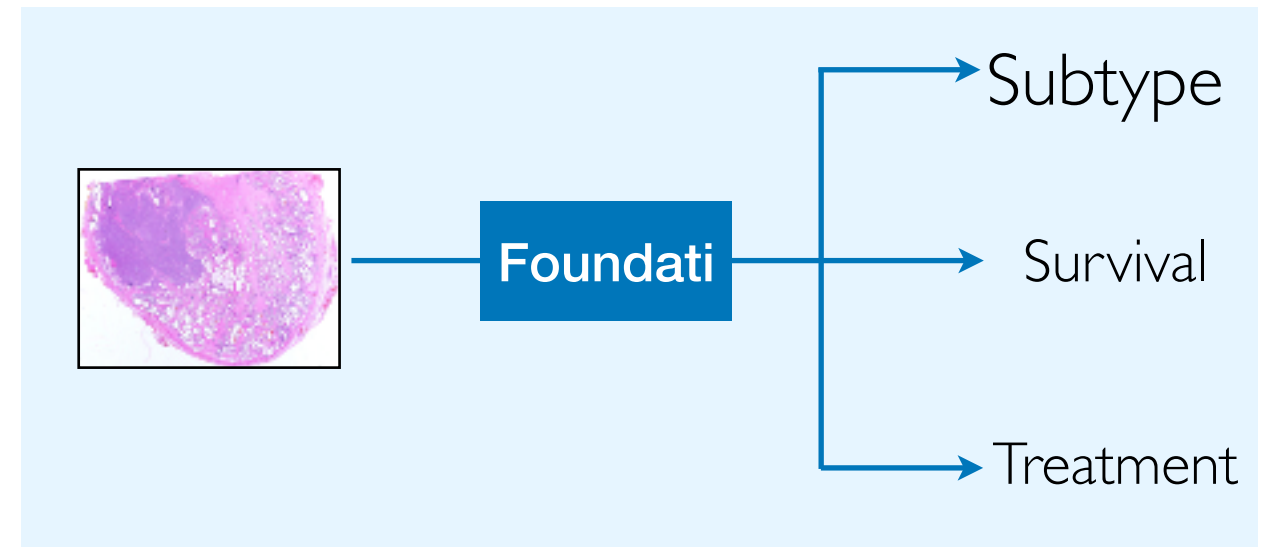


# Today's talk: 3 parts

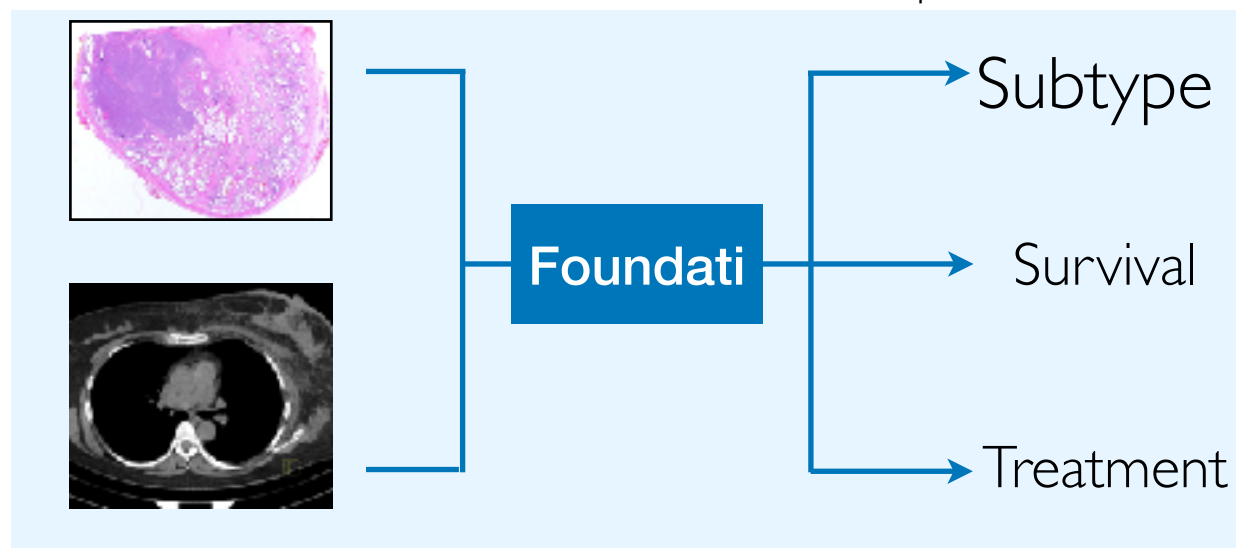
1. Pathology foundation model
2. 3D retinal foundation model
3. A multi-modal foundation model integrating 9 imaging modalities



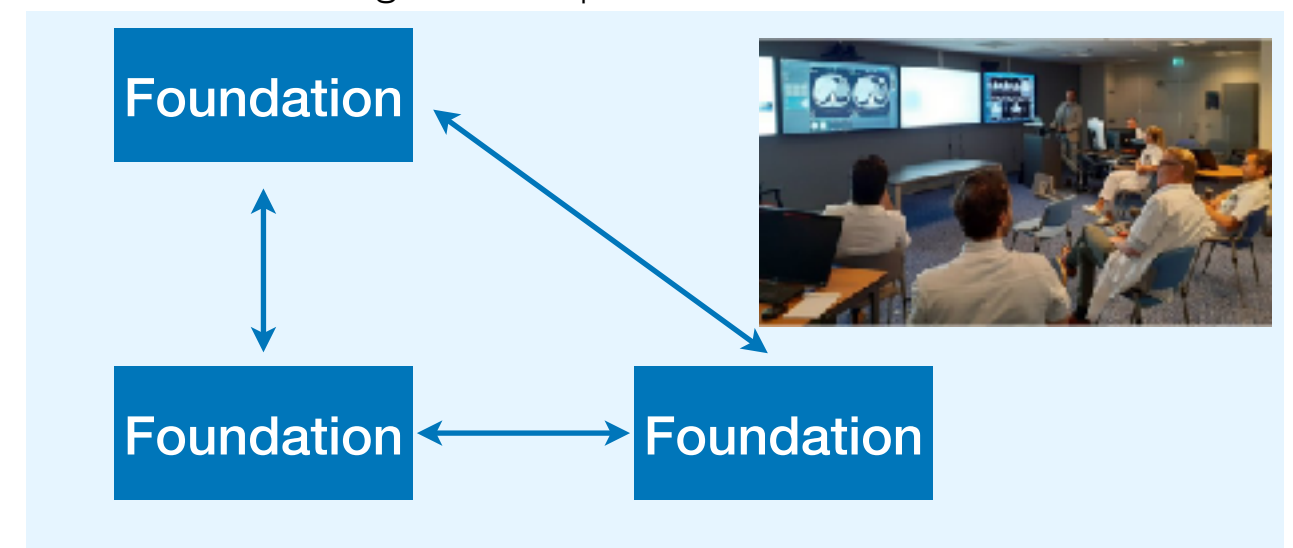
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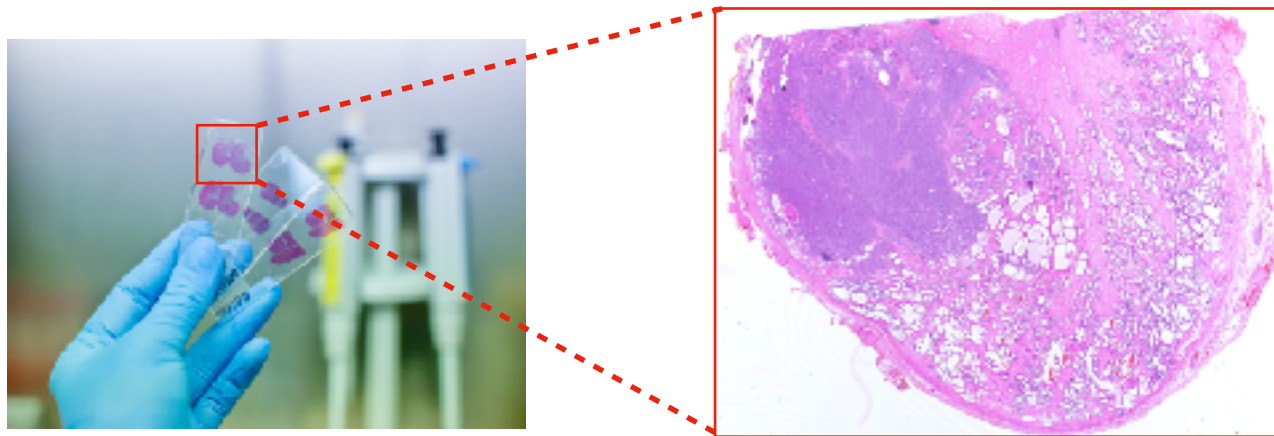
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Integrate multiple foundation models

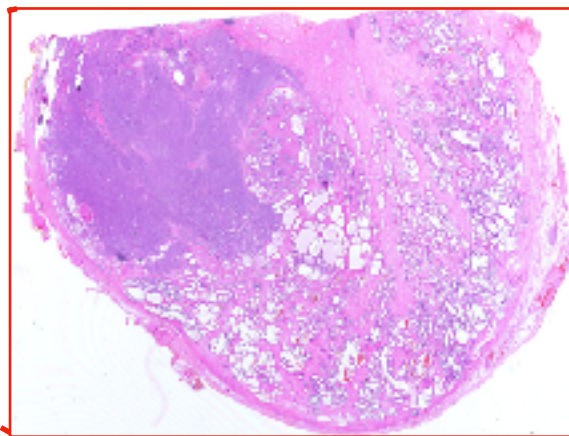
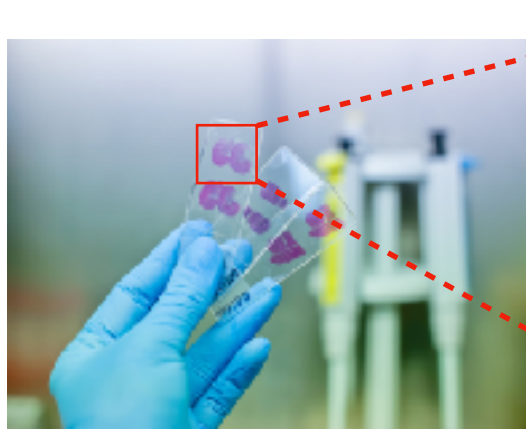


# Pathology images are too large for existing AI models



Pathology images  
100k by 100k pixels

# Pathology images are too large for existing AI models



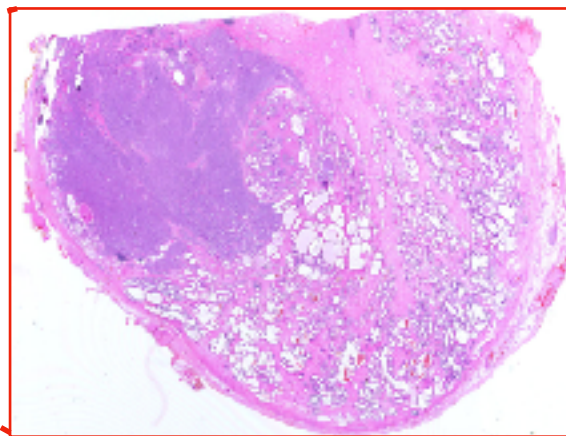
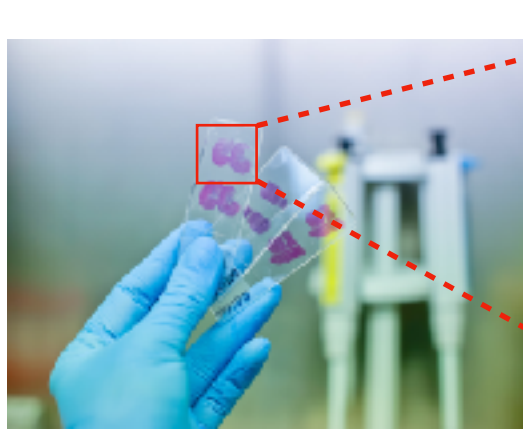
Pathology images  
100k by 100k pixels

As large as a tennis court





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Pathology images  
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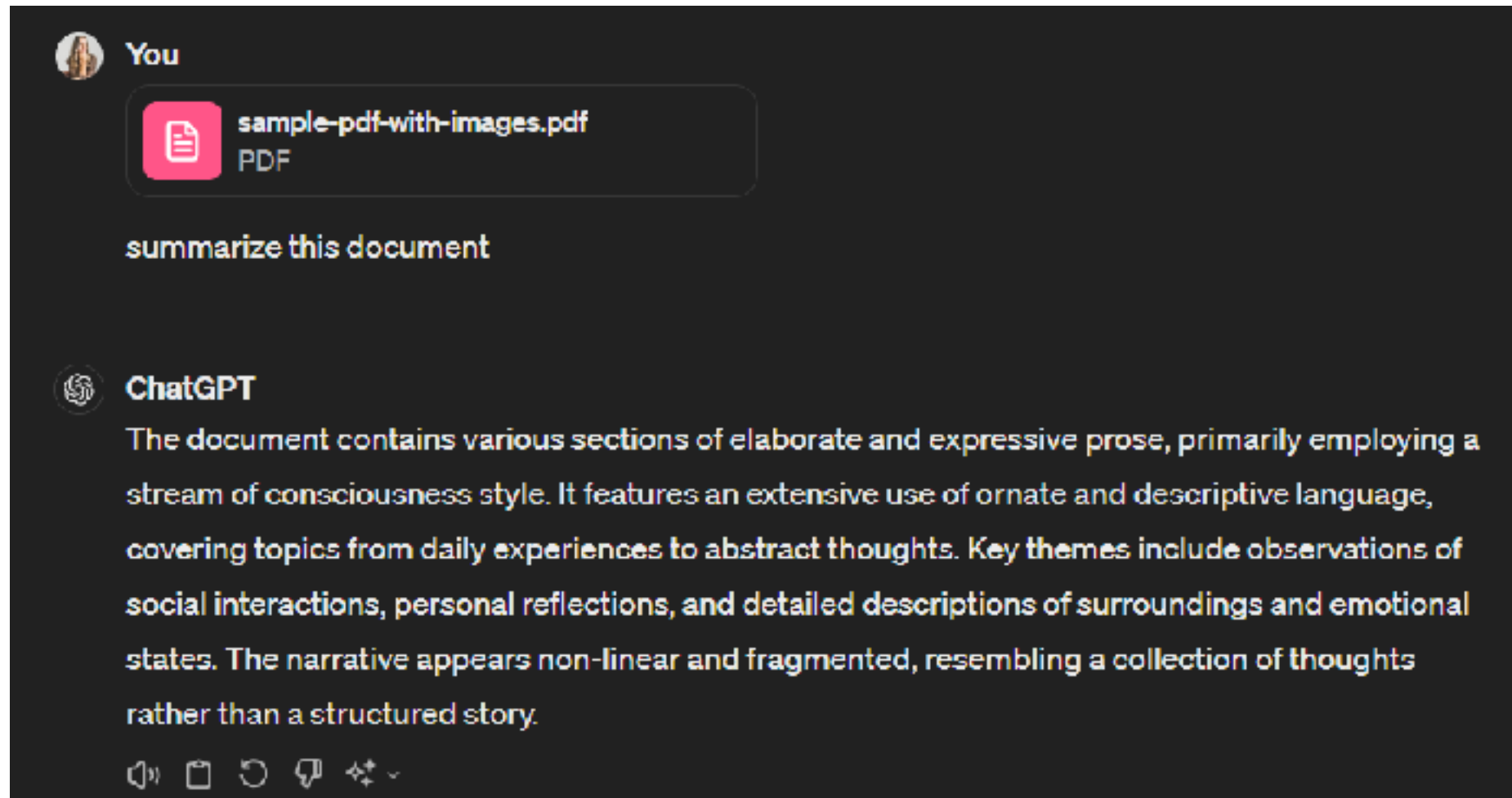


Images handled by existing AI models  
256 by 256 pixels

As large as a tennis ball

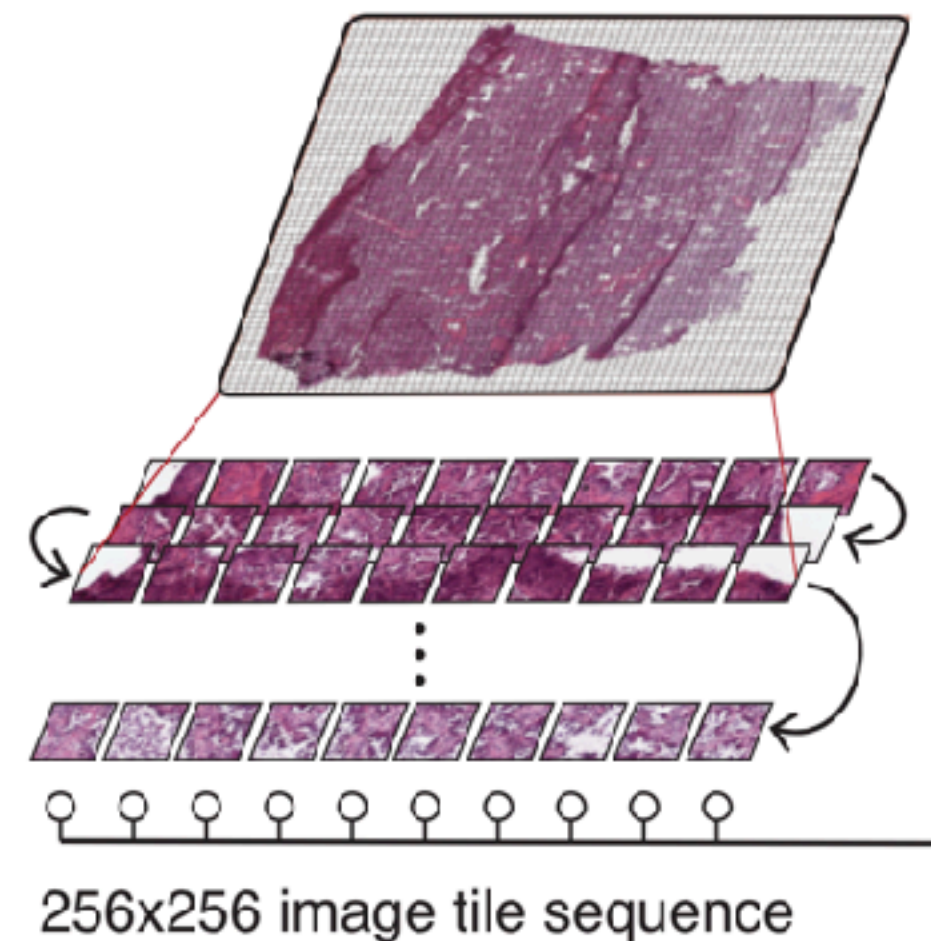
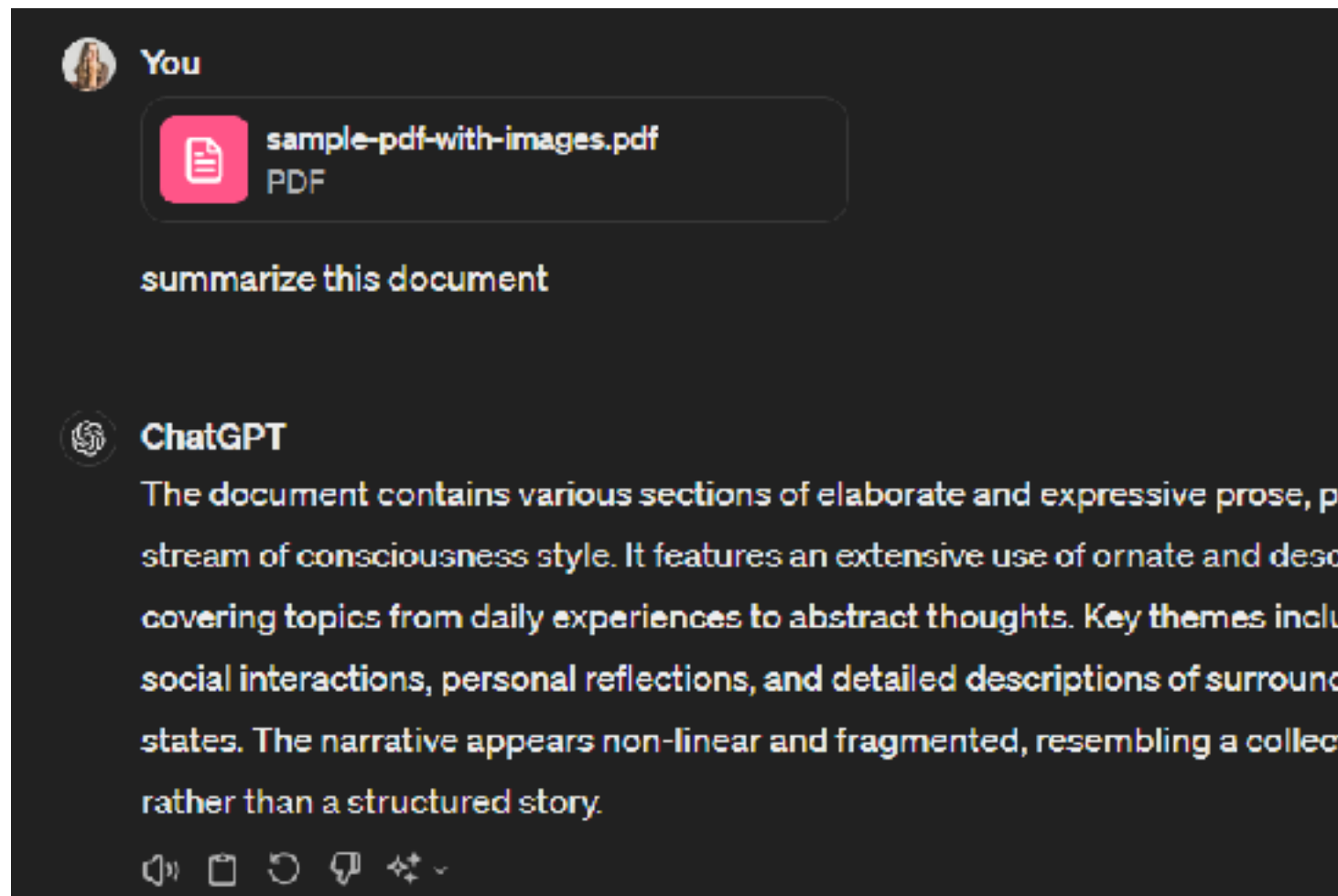


## Long-context modeling: ChatGPT is good at understanding long documents





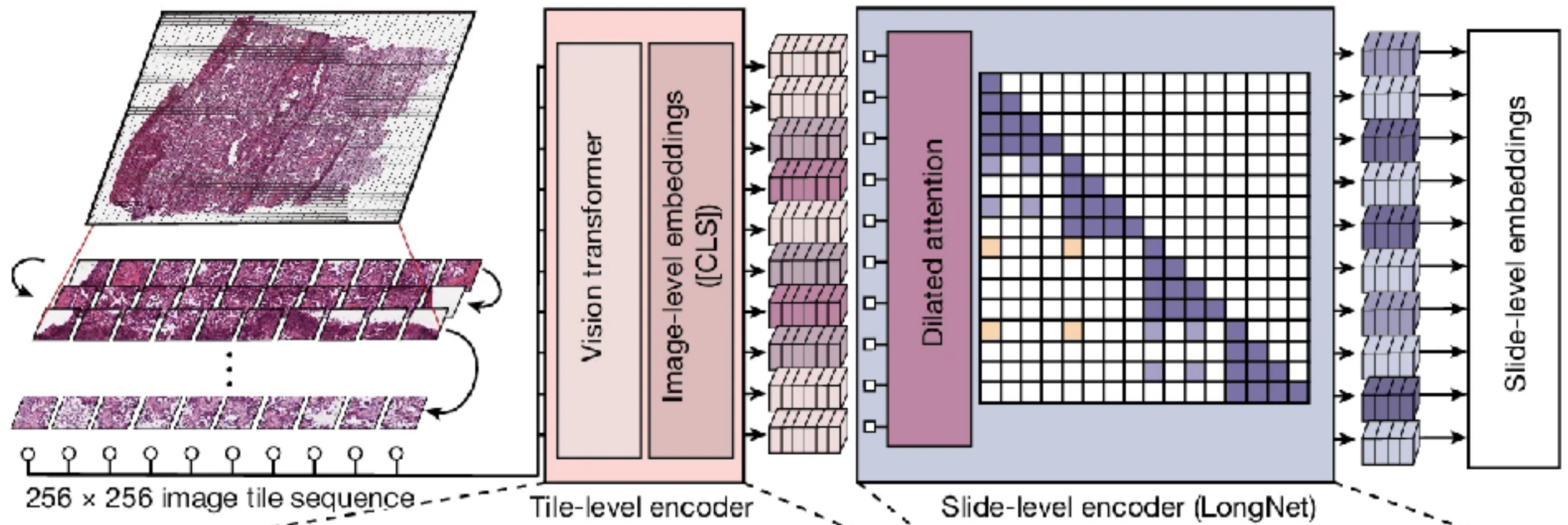
## Long-context modeling: ChatGPT is good at understanding long documents



Use LongNet to model a **long sentence** of small images from a pathology slide

**Tile-level encoder** to capture local patterns

**Slide-level encoder** to capture the pattern in the whole slide



A self-supervised learning framework based on DinoV2 and LongNet

# GigaPath: A whole-slide foundation model for pathology

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Article | [Open access](#) | Published: 22 May 2024

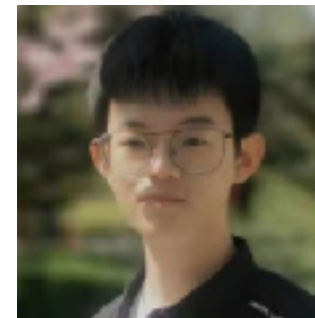
## A whole-slide foundation model for digital pathology from real-world data

Hanwen Xu, Naoto Usuyama, Jaspreet Bagga, Sheng Zhang, Rajesh Rao, Tristan Naumann, Cliff Wong, Zelalem Gero, Javier González, Yu Gu, Yanbo Xu, Mu Wei, Wenhui Wang, Shuming Ma, Furu Wei, Jianwei Yang, Chunyuan Li, Jianfeng Gao, Jaylen Rosemon, Tucker Bower, Scohee Lee, Roshanthi Weerasinghe, Bill J. Wright, Ari Robicsek, Brian Piening, Carlo Bifulco , Sheng Wang  & Hoifung Poon  — [Show fewer authors](#)

[Nature](#) **630**, 181–188 (2024) | [Cite this article](#)

200k model downloads every month  
(Hugging Face)!

Media coverage: [Forbes](#), [Yahoo](#), [Becker's hospital review](#), [Fierce biotech](#), [CTOL digital solutions](#), [HIT consultant](#), [GeekWire](#), [Cosmic log](#), [HealthXL](#), [RamaOnHealthcare](#), [Providence](#), [nikkei](#), [cryptorank](#), [deeptech](#)



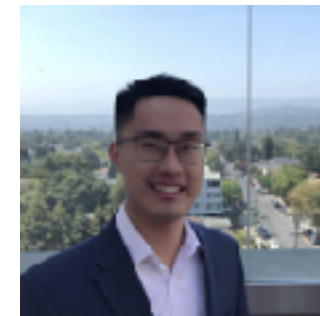
Hanwen Xu  
U of Washington



Naoto Usuyama  
Microsoft Research



Carlo Bifulco  
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Sheng Wang  
U of Washington

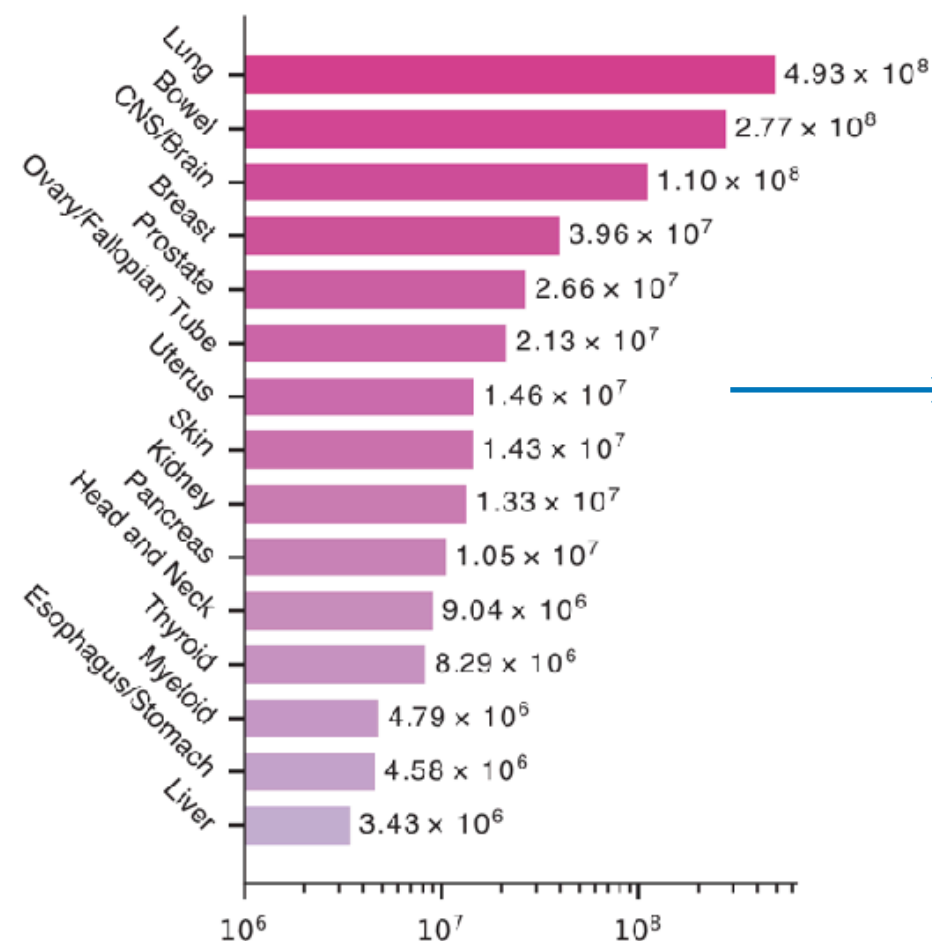


Hoifung Poon  
Microsoft Research



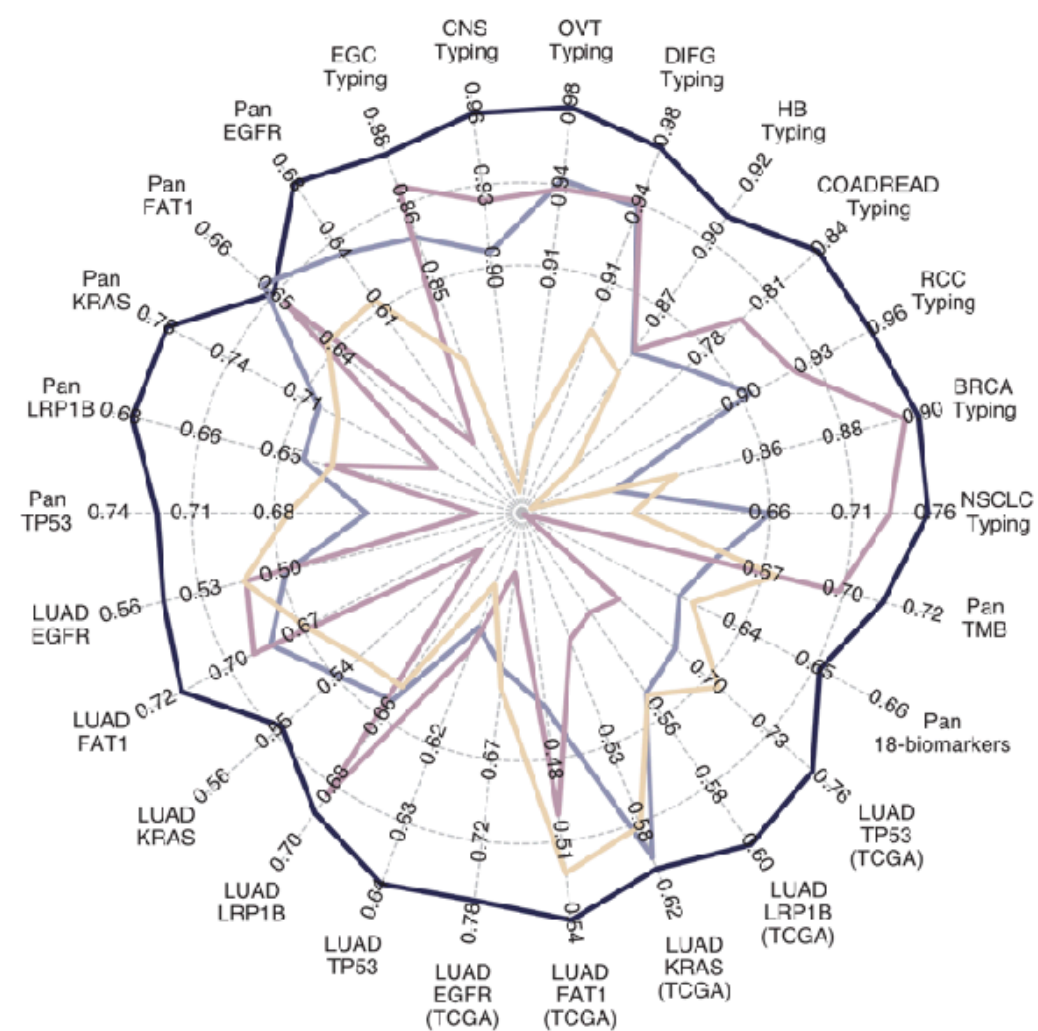
# One (foundation) model fits all

30,000 patients from 28 hospitals  
140k slides, 15 cancer types



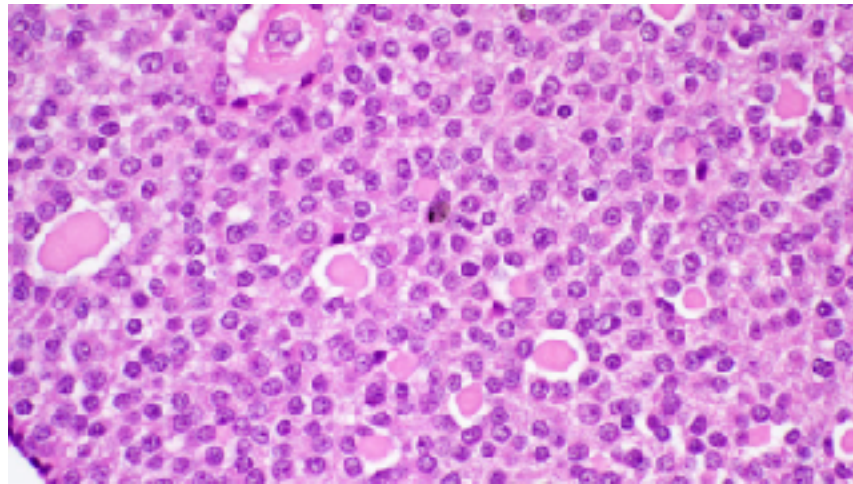
GigaPa

Best performance on 25 out of 26 tasks



# Cancer subtype classification using pathology images

## Ovarian



?

EOV

CCO

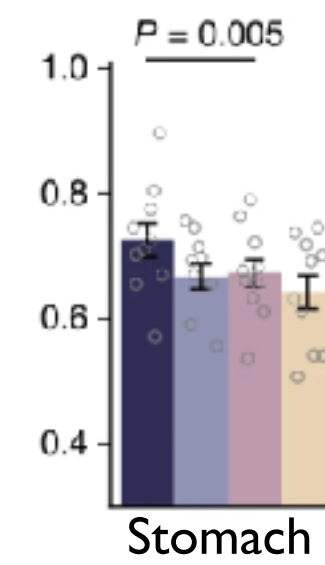
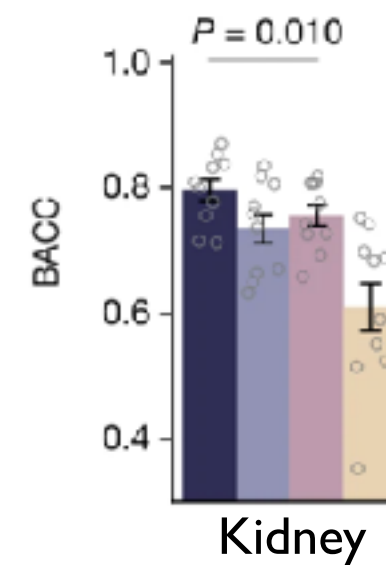
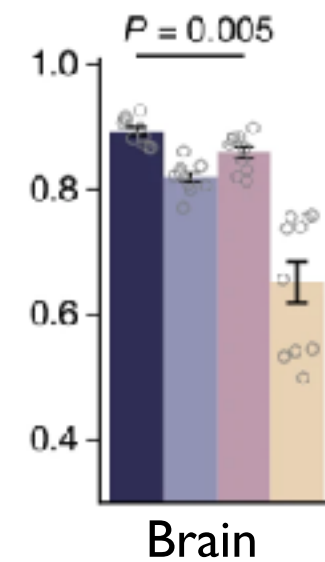
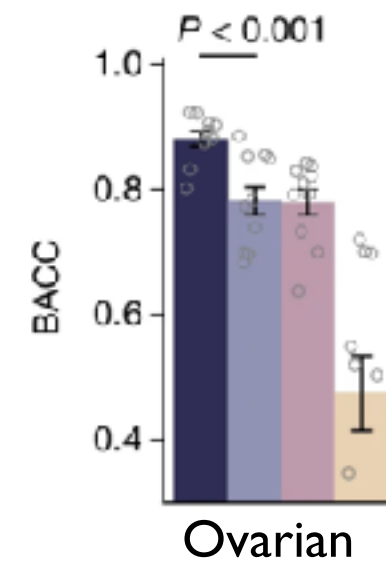
HGSO

LGSCO

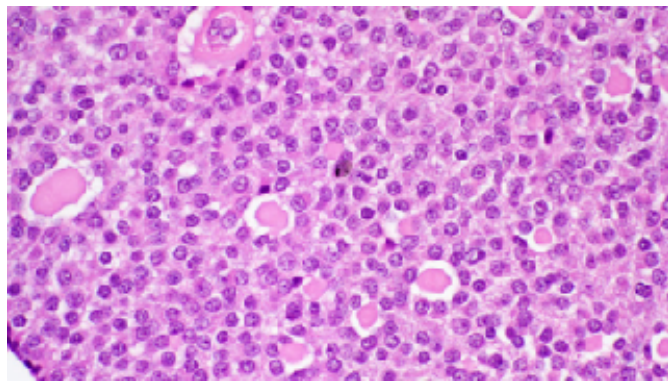
MO

OC

■ GigaPath ■ HIPT ■ CtransPath ■ REMEDIS



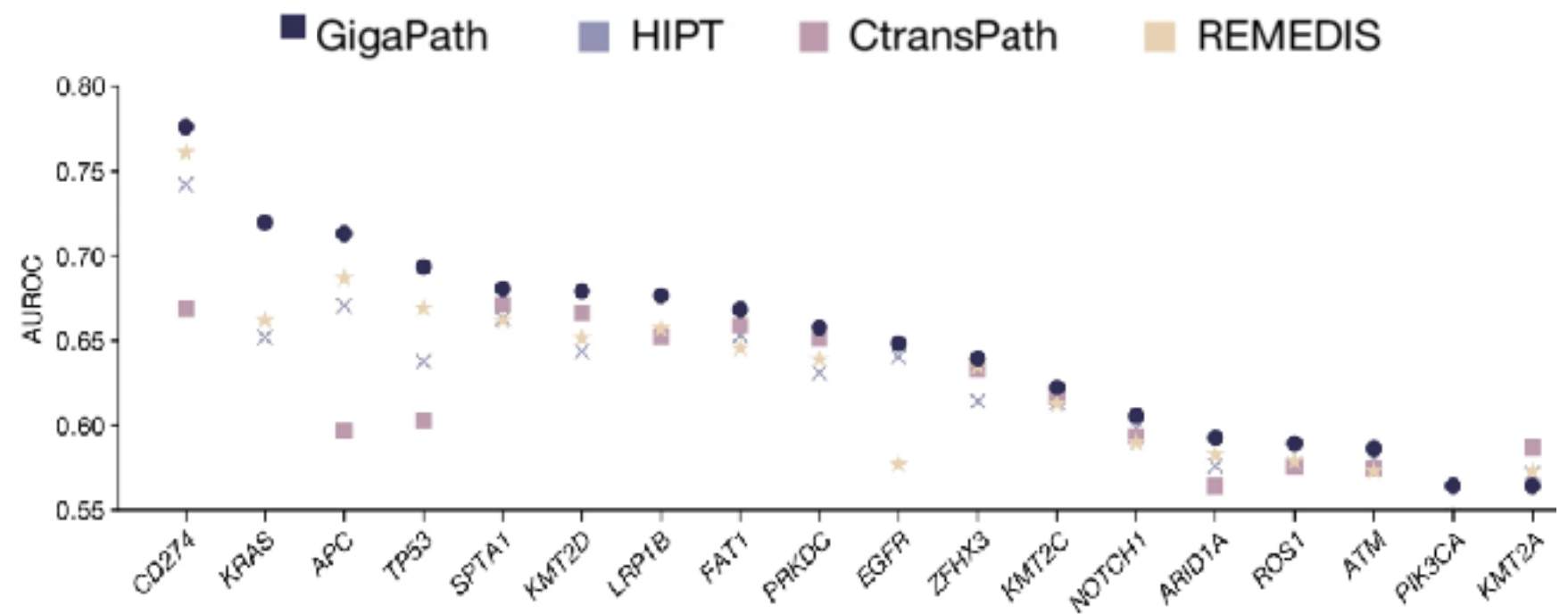
# Biomarker prediction using pathology images: best performance on 17 out of 18 biomarker predictions



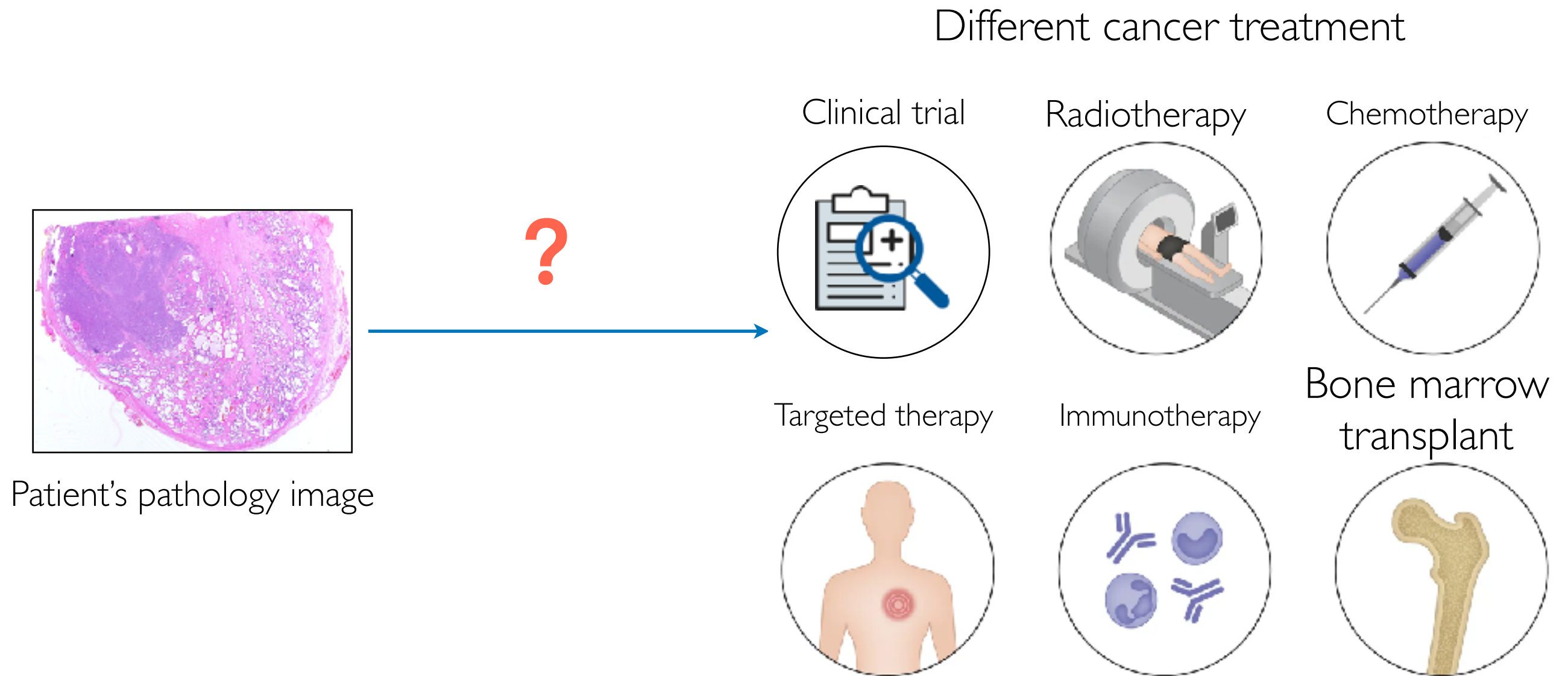
KRAS?

Mutate

Non-



# Directly predicting treatment is too difficult





# Chain-of-Thought: Decompose a complicated task into many subtasks

## Standard GenAI

### Model Input

Q: Roger has 5 tennis balls. He buys 2 more cans of tennis balls. Each can has 3 tennis balls. How many tennis balls does he have now?

A: The answer is 11.

Q: The cafeteria had 23 apples. If they used 20 to make lunch and bought 6 more, how many apples do they have?

### Model Output

A: The answer is 27. ❌

## Chain-of-Thought GenAI

### Model Input

Q: Roger has 5 tennis balls. He buys 2 more cans of tennis balls. Each can has 3 tennis balls. How many tennis balls does he have now?

A: Roger started with 5 balls. 2 cans of 3 tennis balls each is 6 tennis balls.  $5 + 6 = 11$ . The answer is 11.

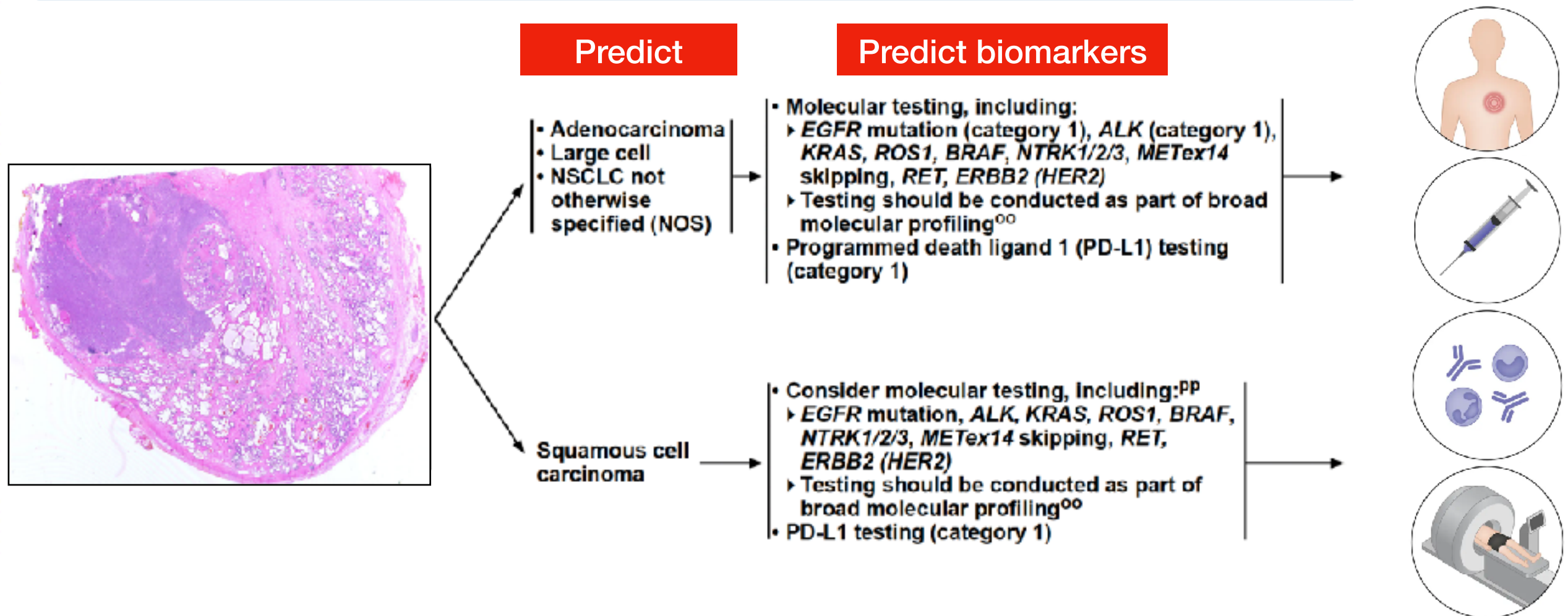
Q: The cafeteria had 23 apples. If they used 20 to make lunch and bought 6 more, how many apples do they have?

### Model Output

A: The cafeteria had 23 apples originally. They used 20 to make lunch. So they had  $23 - 20 = 3$ . They bought 6 more apples, so they have  $3 + 6 = 9$ . The answer is 9. ✅



# Our idea: use Clinical Guideline as the Chain-of-Thought

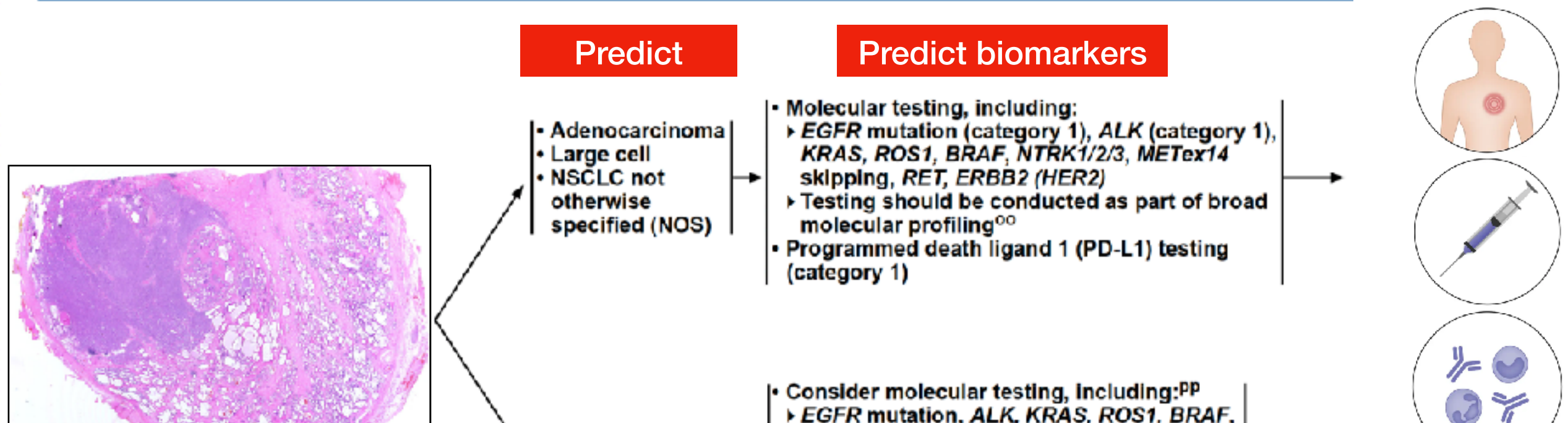


# Our idea: use Clinical Guideline as the Chain-of-Thought



National  
Comprehensive  
Cancer  
Network®

## NCCN Guidelines Version 7.2024 Non-Small Cell Lung Cancer



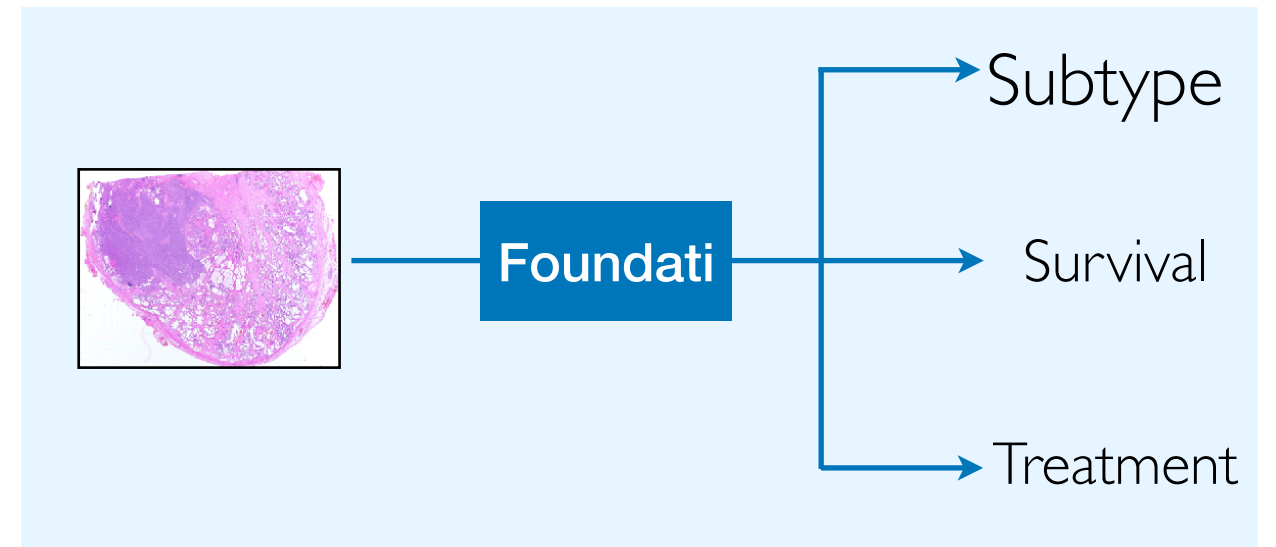
Human-AI collaboration: Experts derive the guideline. AI makes decision on each branch.  
Future implication: AI model as a clinical lab test

# Today's talk: 3 parts

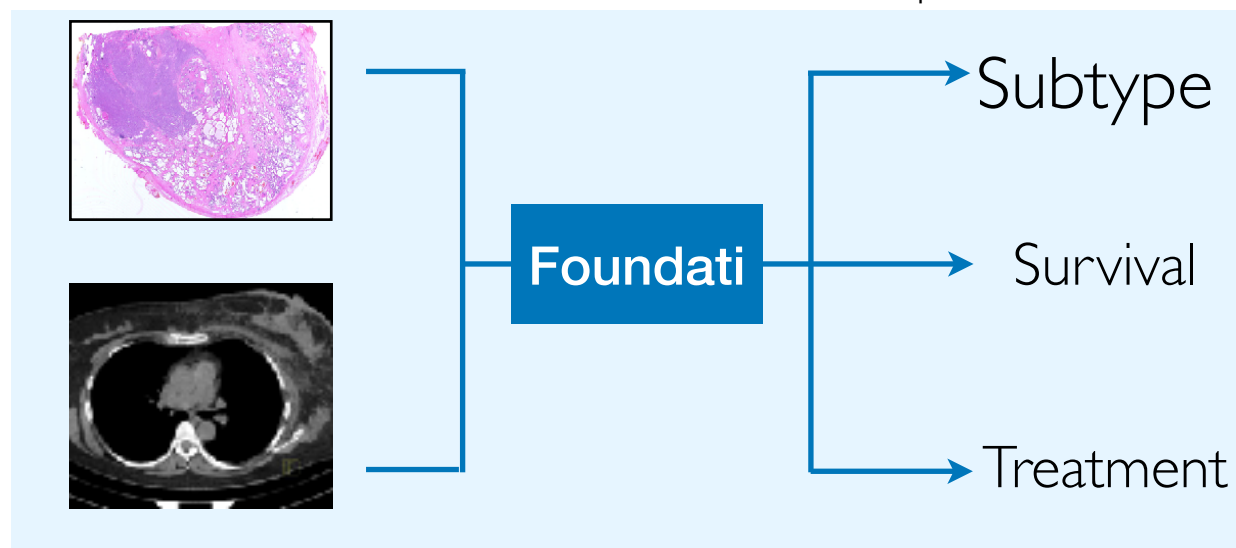
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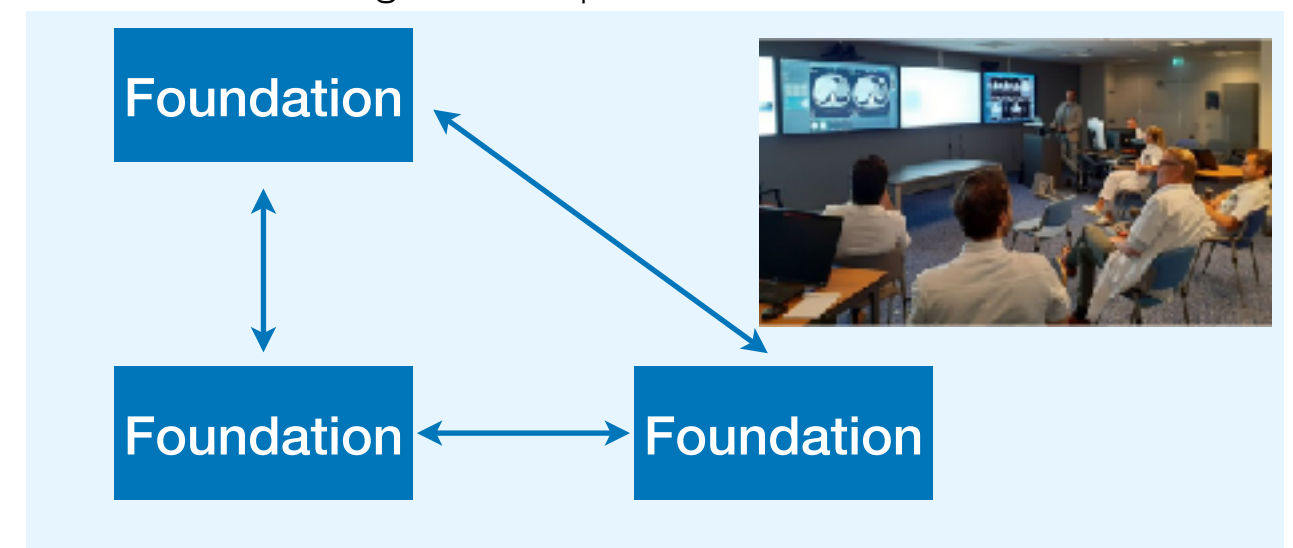
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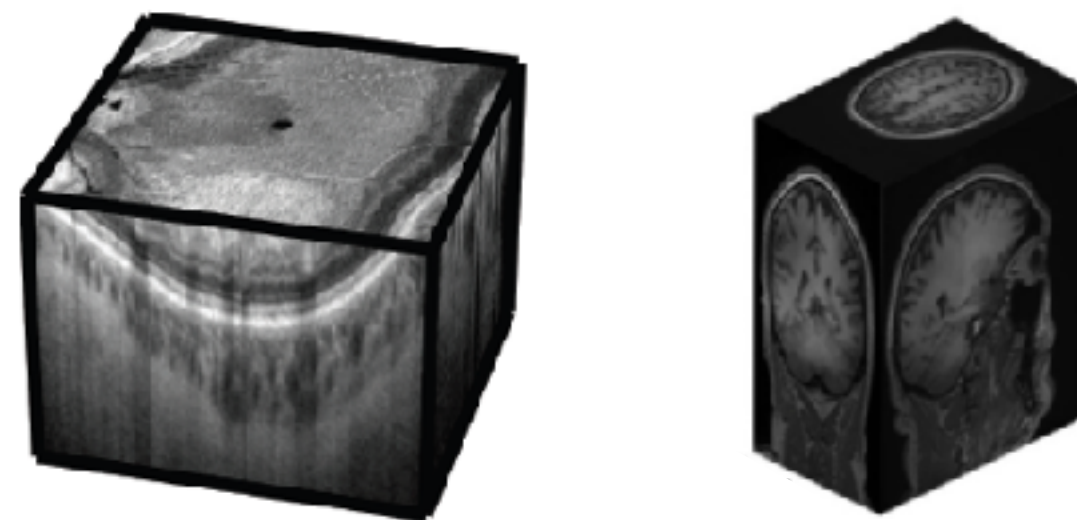




## Lots of medical images are 3D



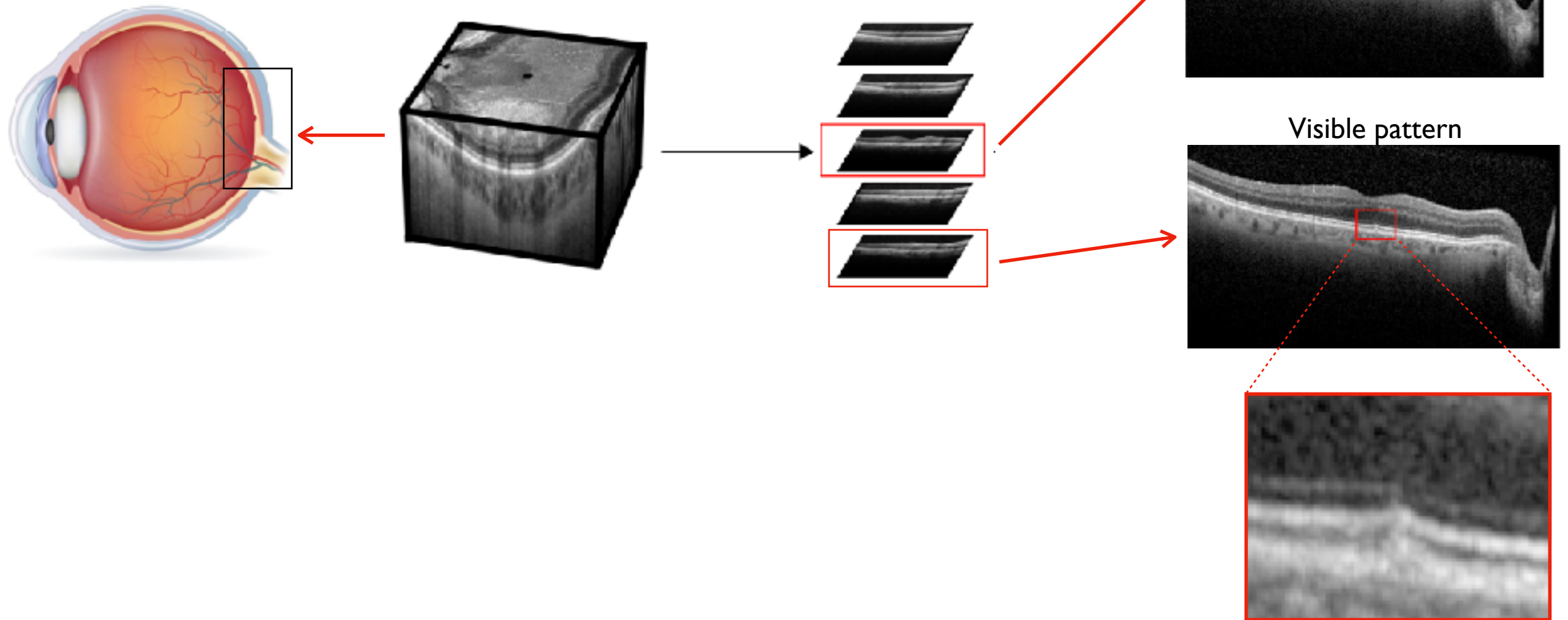
General-domain images: 2D



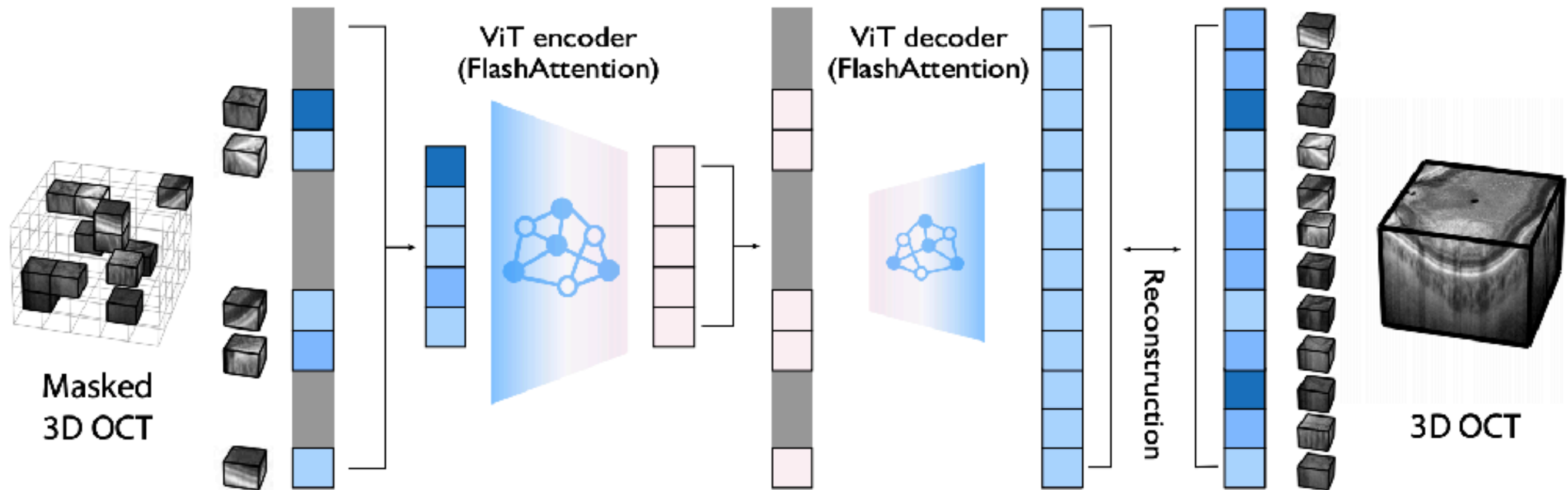
3D biomedical images: CT, MRI, OCT

# Disease pattern is very small and hard to detect

Age-related macular degeneration



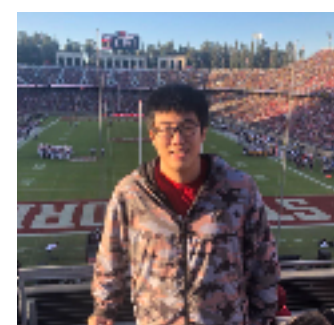
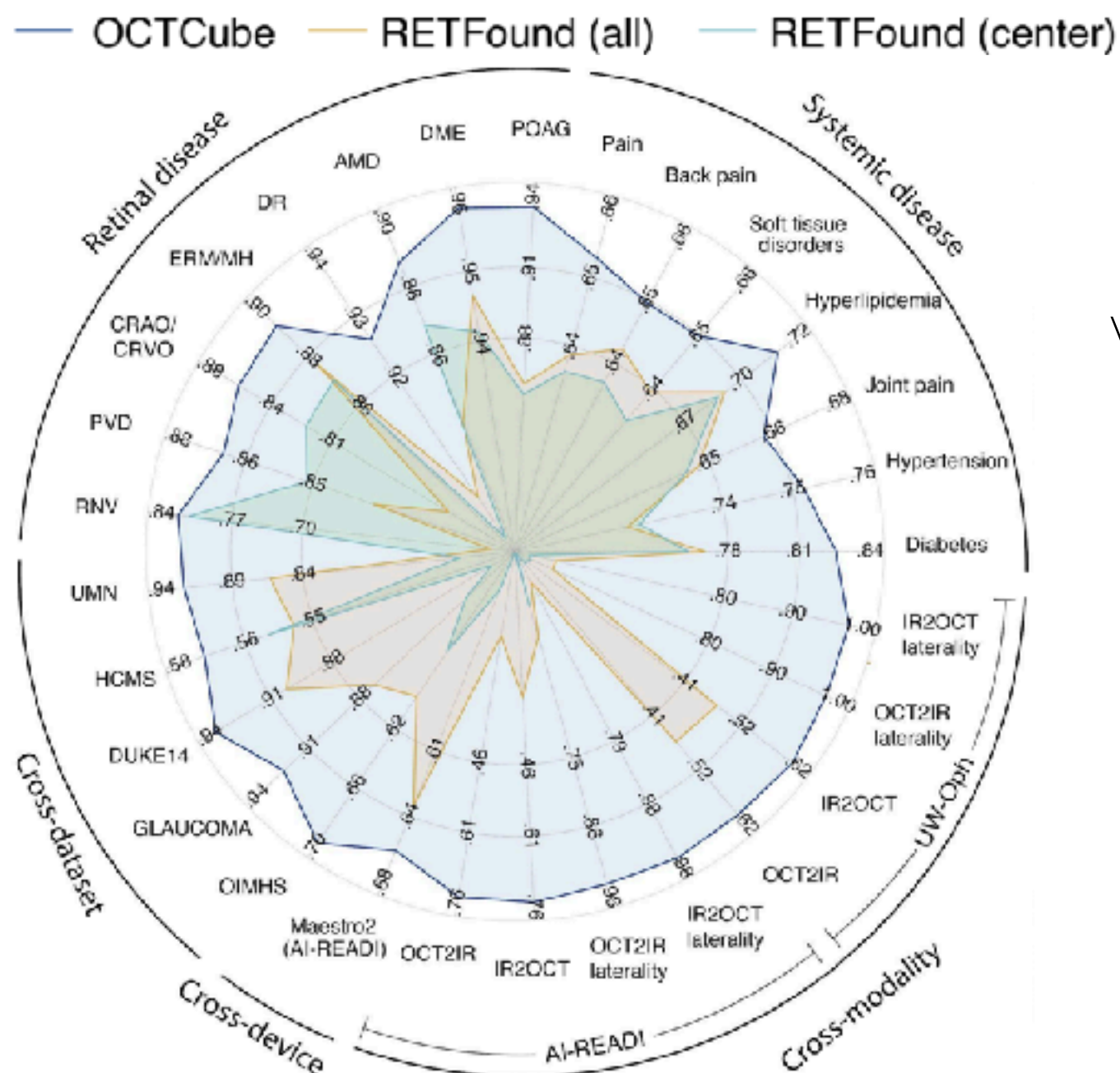
# A GenAI framework for 3D medical imaging: MAE + FlashAttention



Convert a 3D volume to a **long sentence** of small cubes



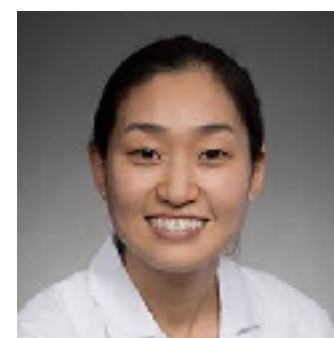
The first 3D OCT foundation model trained from 26,605 patients  
Best performance on all 29 tasks



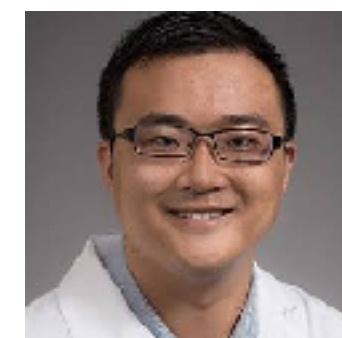
Work done by Zixuan Liu  
U of Washington



Dr. Miao Zhang  
Genentech

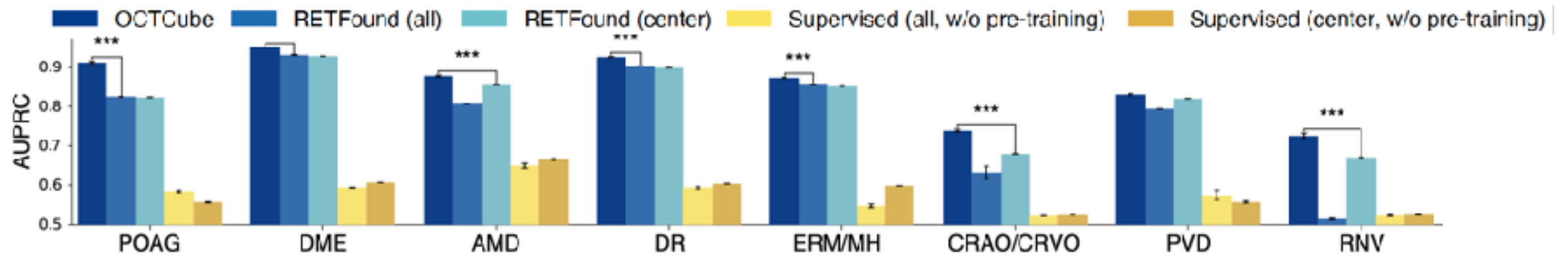


Dr. Cecilia Lee  
UW ophthalmology



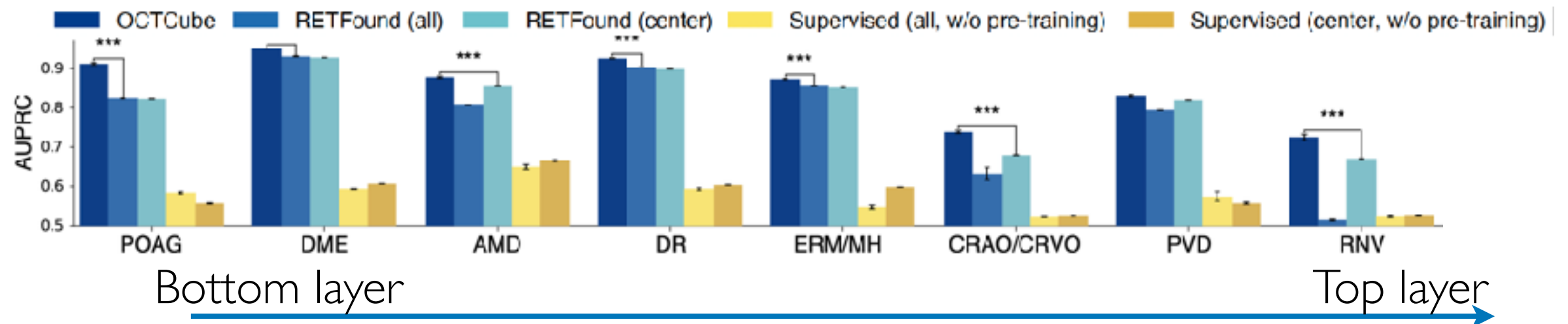
Dr. Aaron Lee  
UW ophthalmology

## 3D model offers accurate and interpretable predictions for retinal diseases

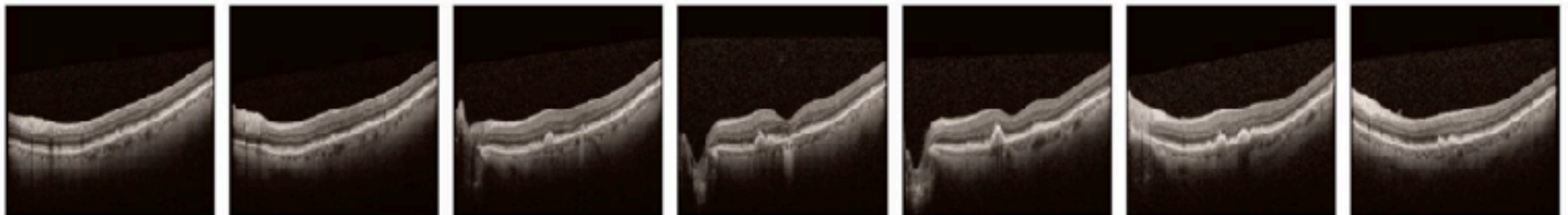




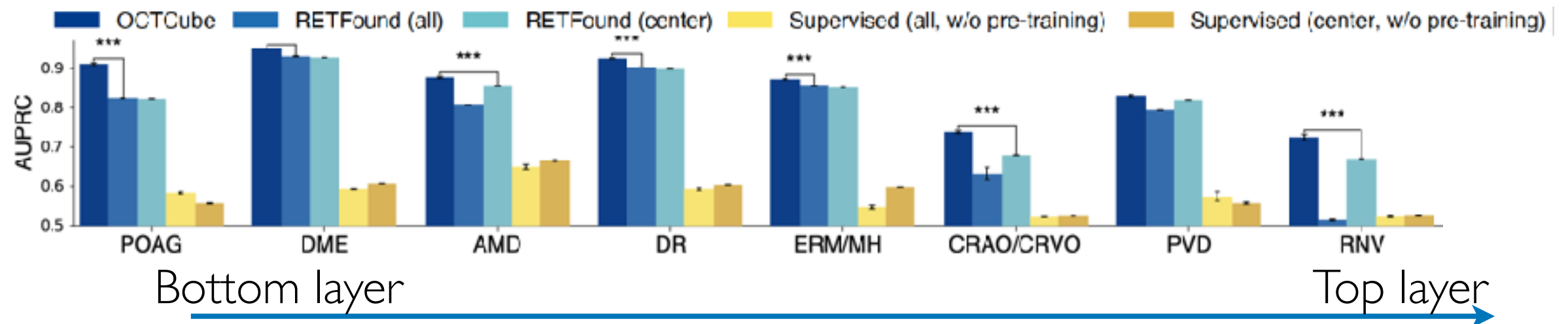
## 3D model offers accurate and interpretable predictions for retinal diseases



OCT slice

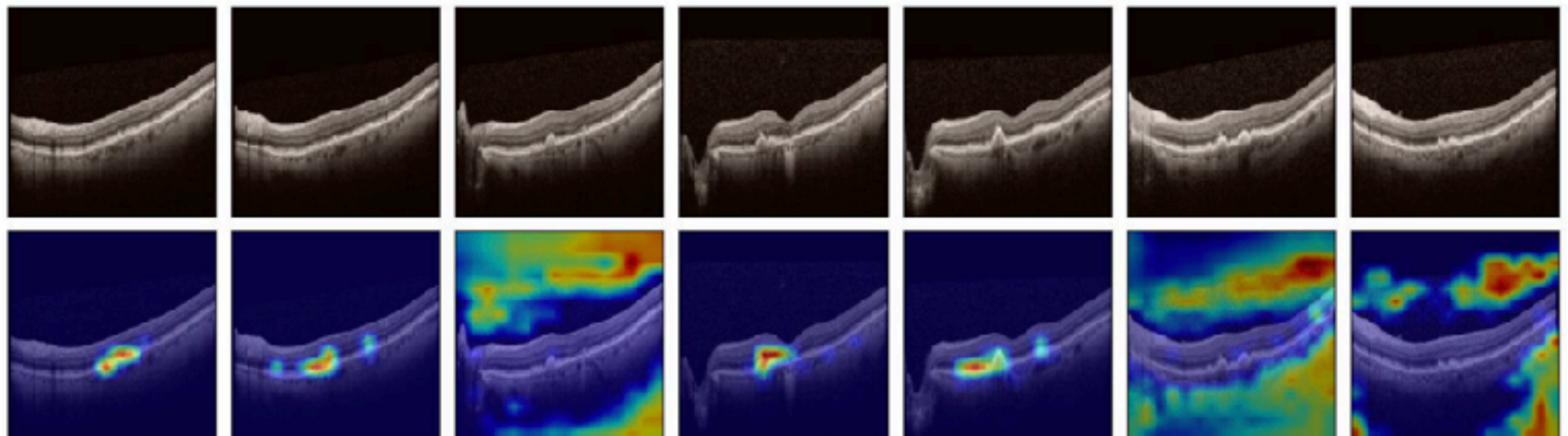


## 3D model offers accurate and interpretable predictions for retinal diseases

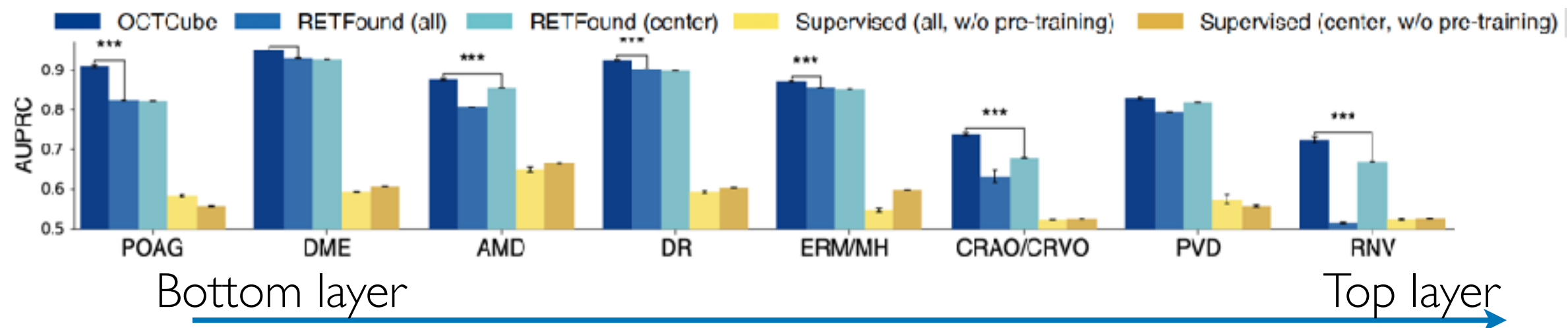


OCT slice

RETFound



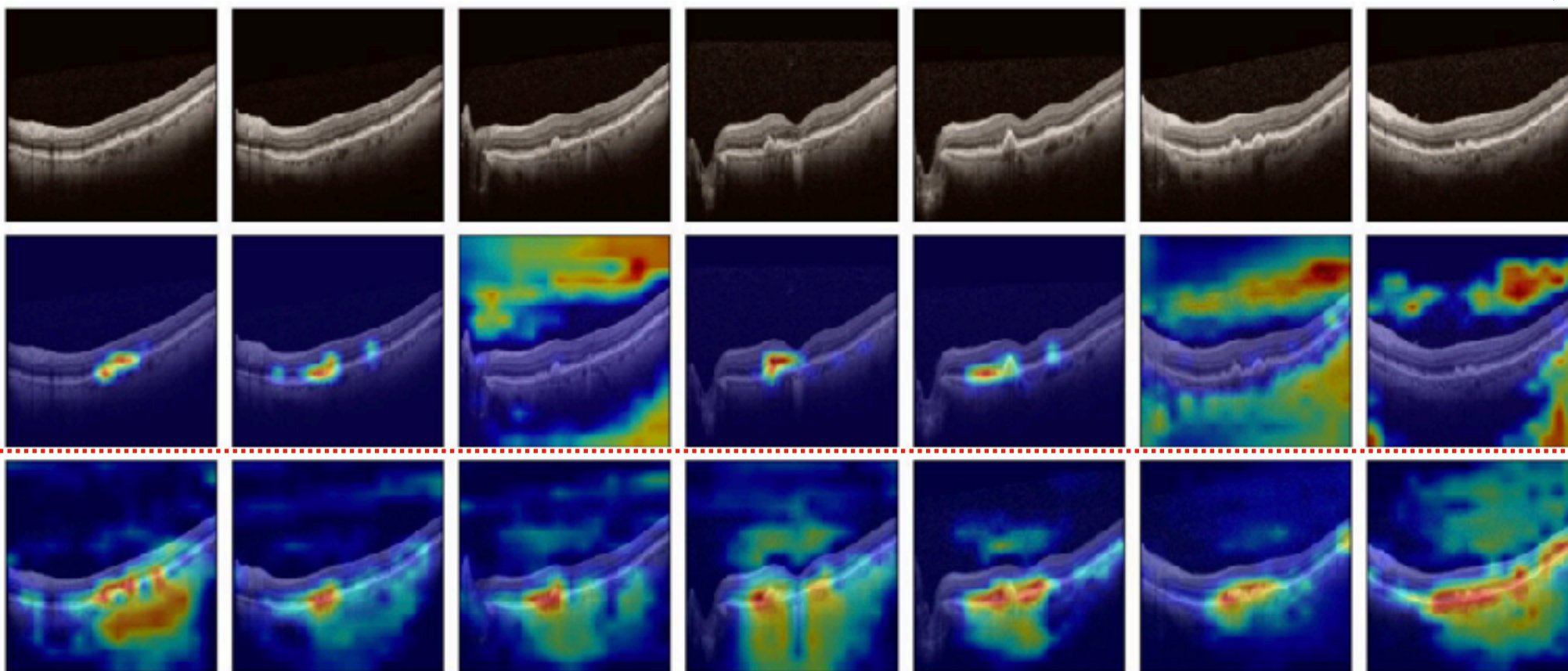
## 3D model offers accurate and interpretable predictions for retinal diseases



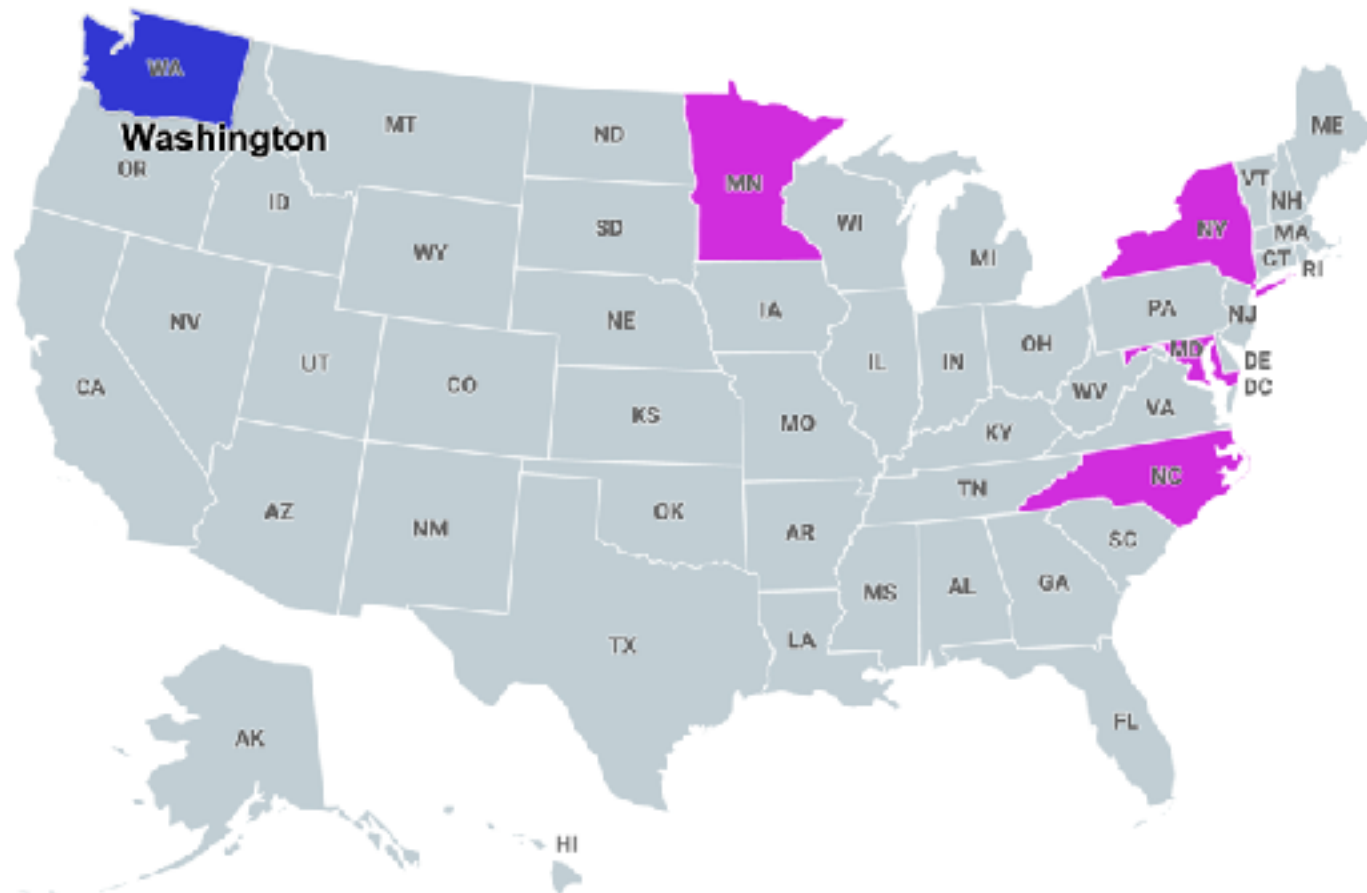
OCT slice

RETFound

Our method



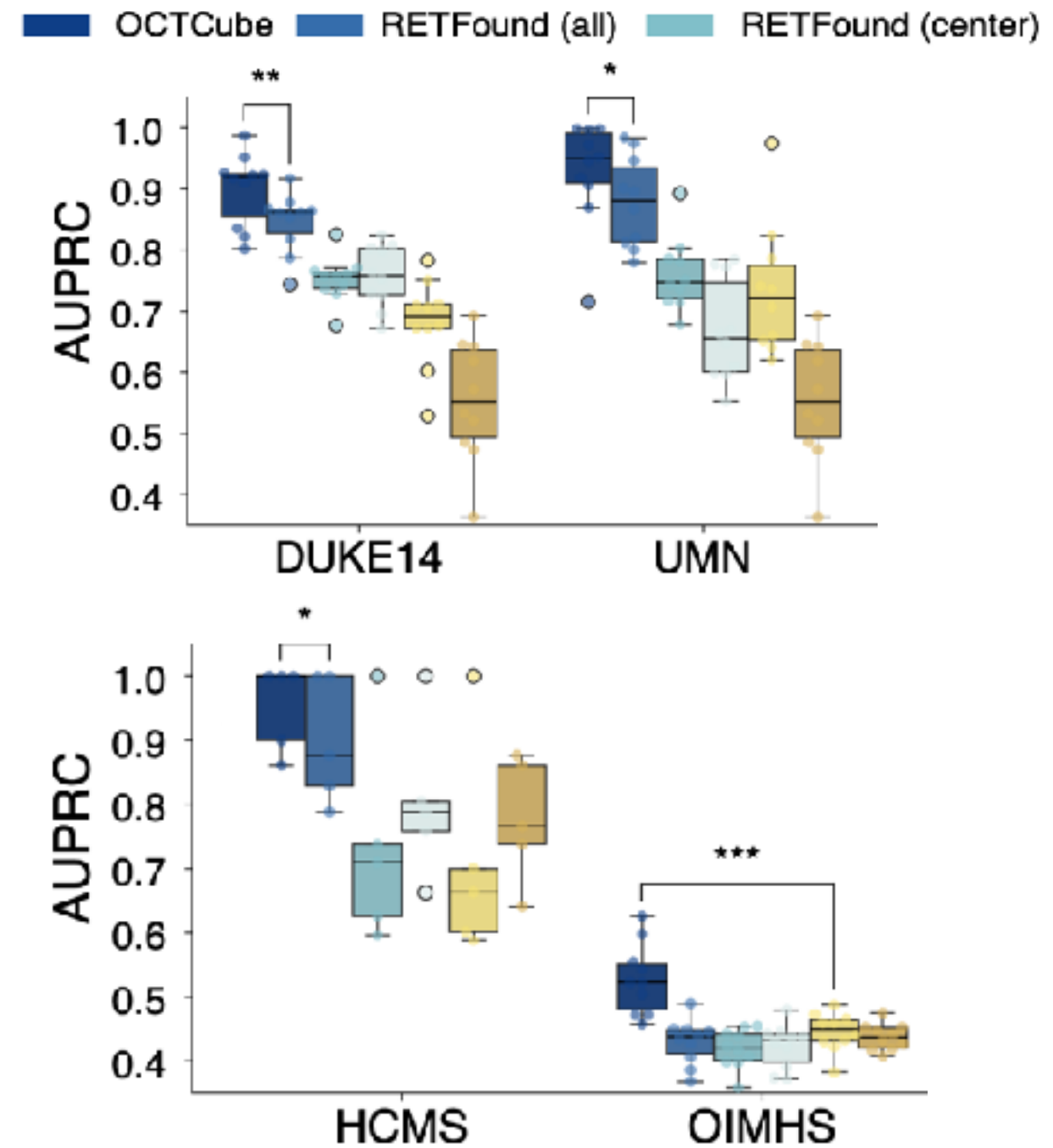
3D model is better at generalization: Cross-cohort prediction at Duke, University of Minnesota, Johns Hopkins, NYU



Training



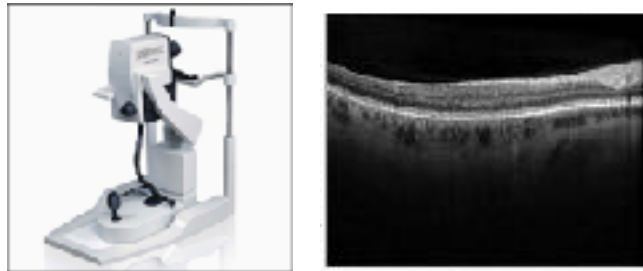
Test



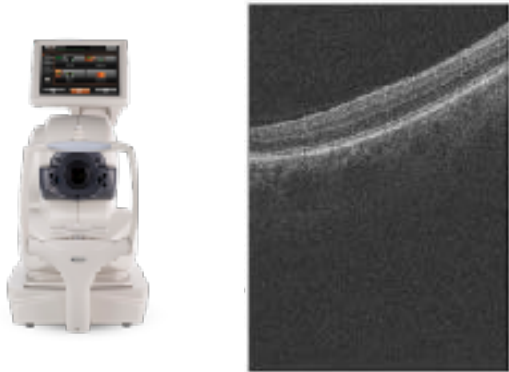


## 3D model enables cross-device prediction

Heidelberg Spectralis



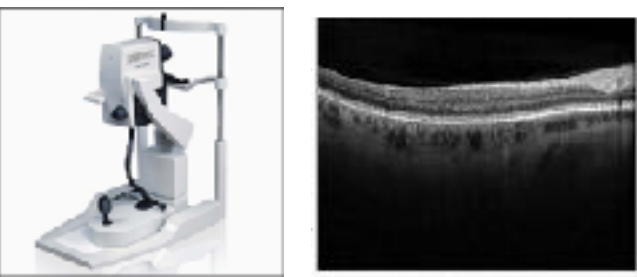
Topcon Masestro2



3D model enables cross-device prediction

OCTCube RETFound (all) RETFound (center)

Heidelberg Spectralis



Pretrai

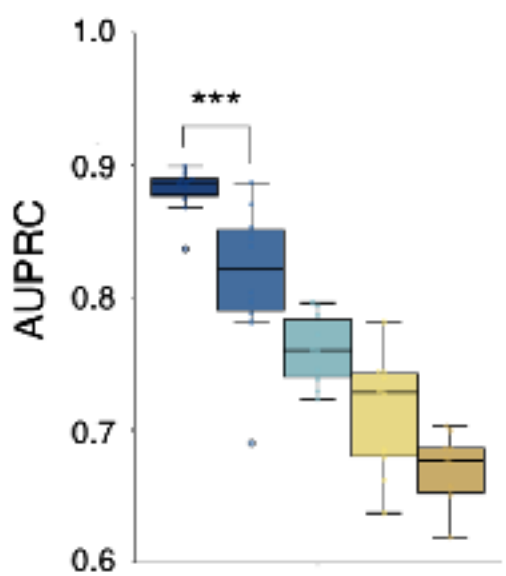


Heidelberg Spectralis

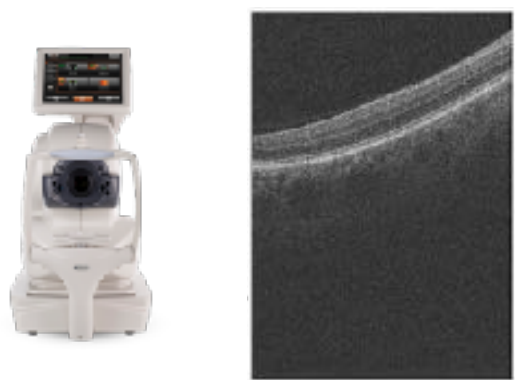
Fine-



Topcon Masestro2



Topcon Masestro2



Pretrai

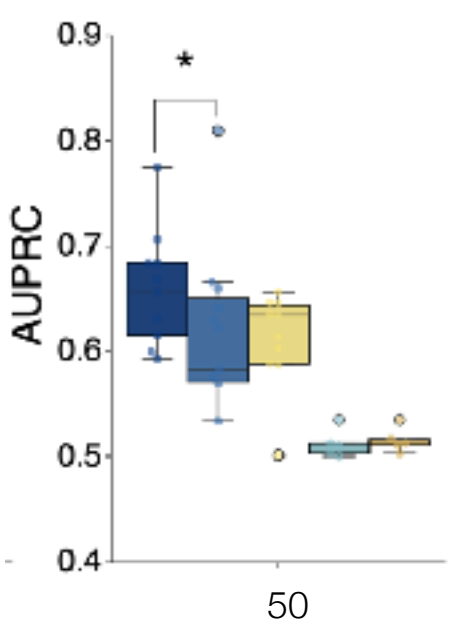


Heidelberg Spectralis

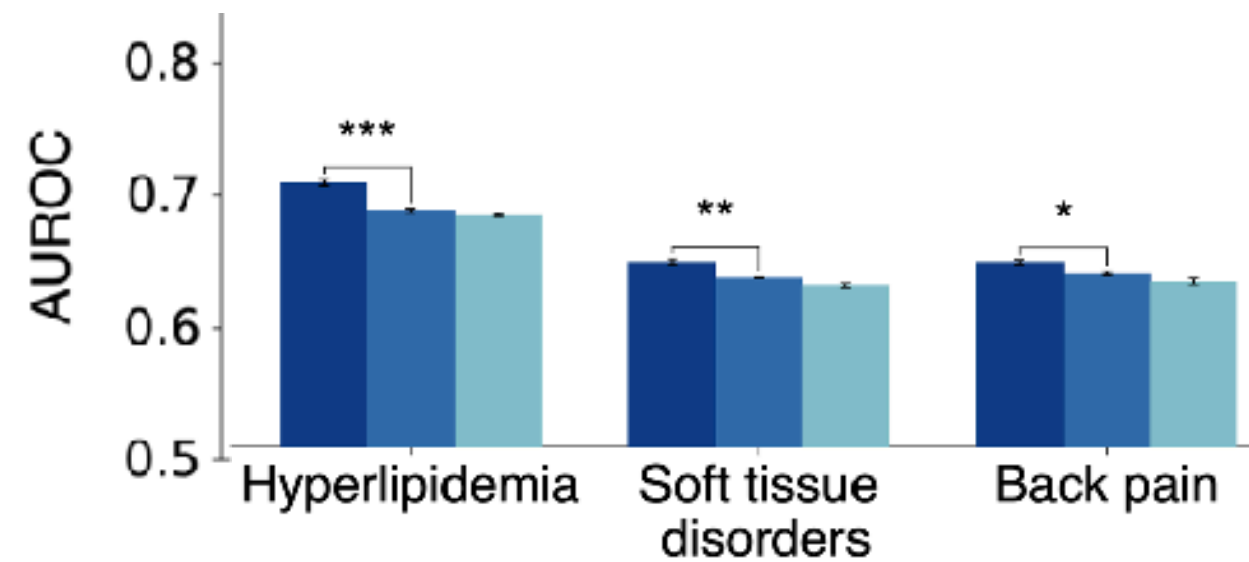
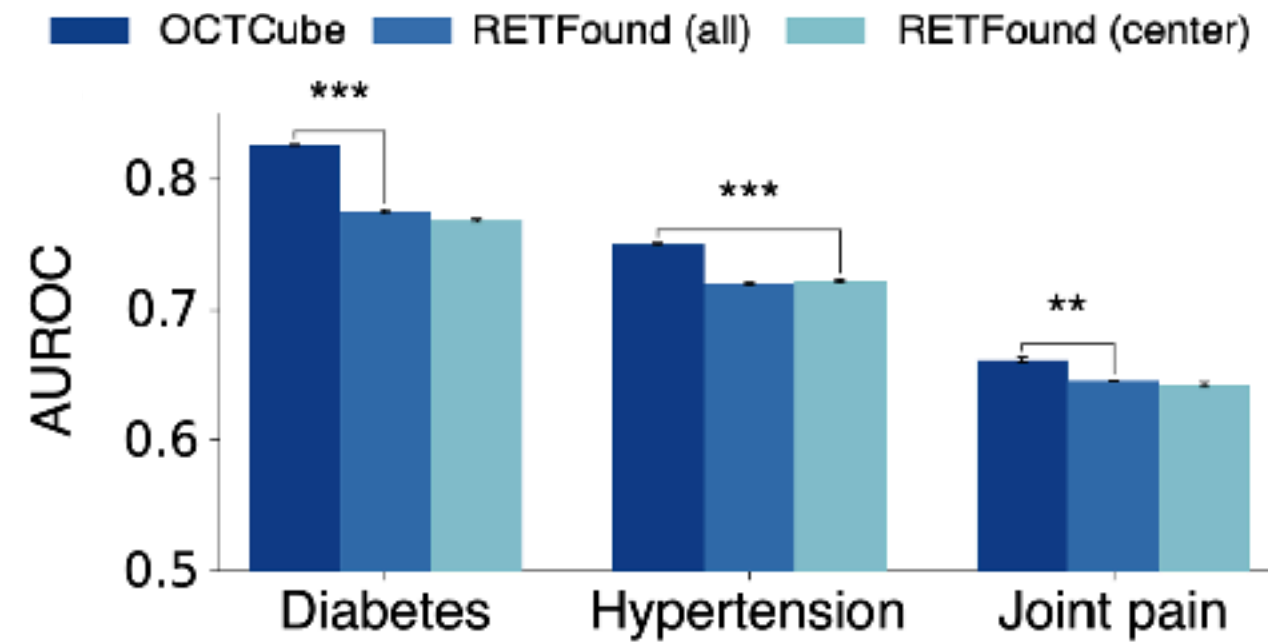
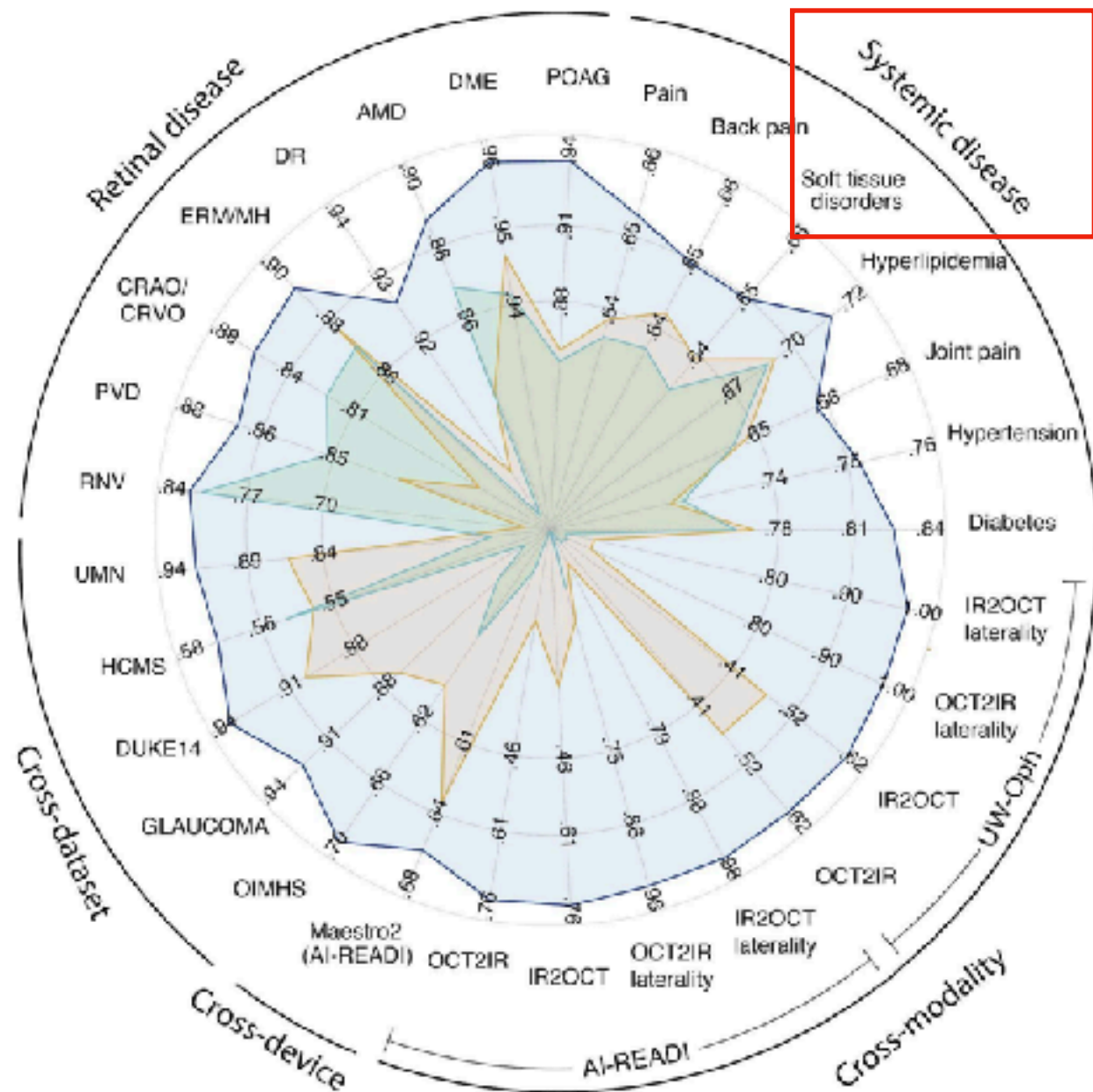
Fine-



Zeiss cirrus

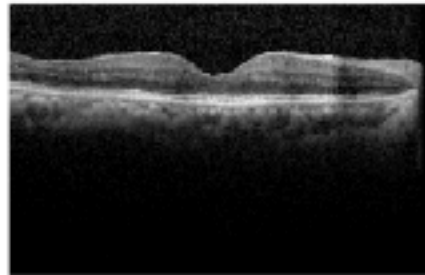


# Predict systemic diseases

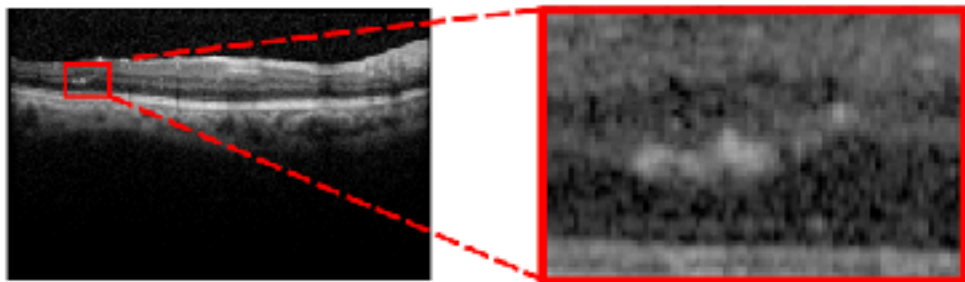
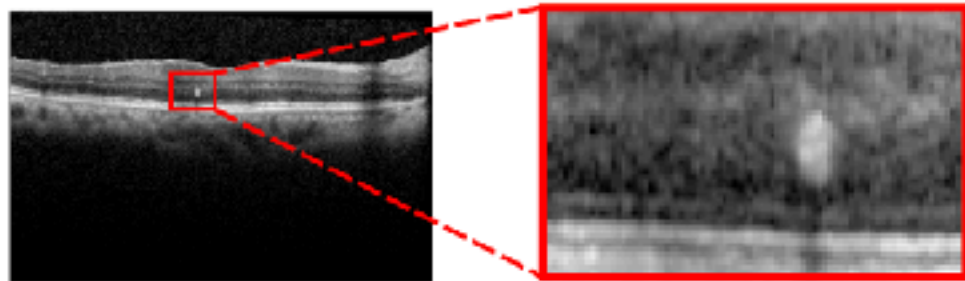


# One-year early prediction of diabetes

1st visit: **2D model** cannot identify diabetes



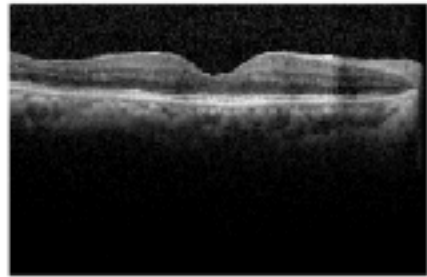
1st visit: **3D model** detects diabetes signal



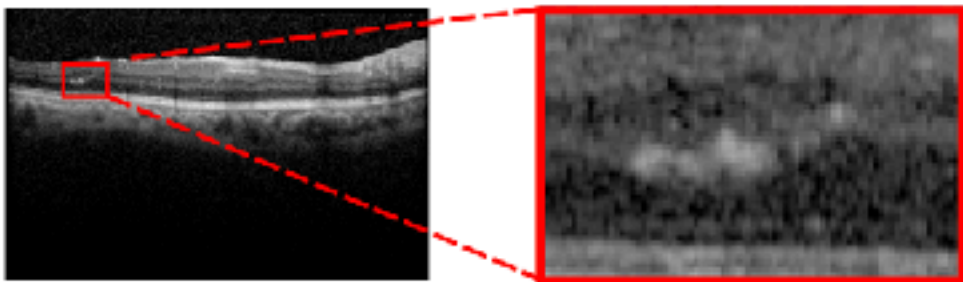
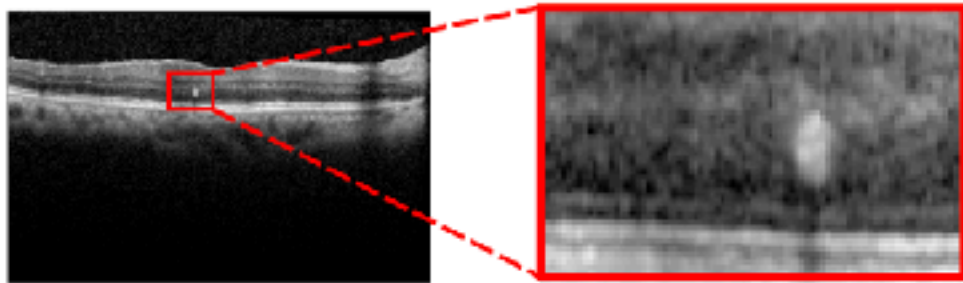


# One-year early prediction of diabetes

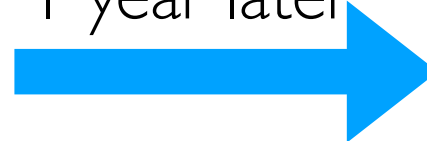
1st visit: **2D model** cannot identify diabetes



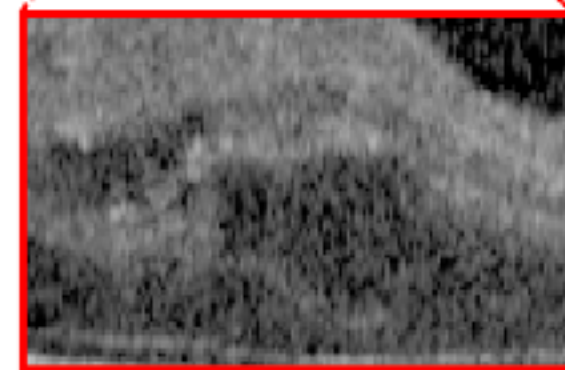
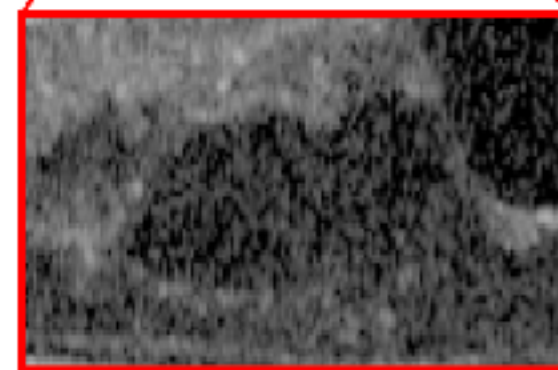
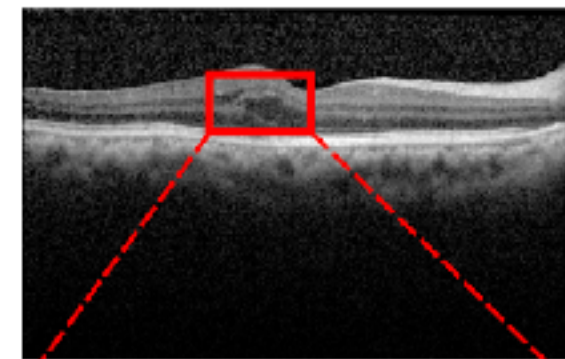
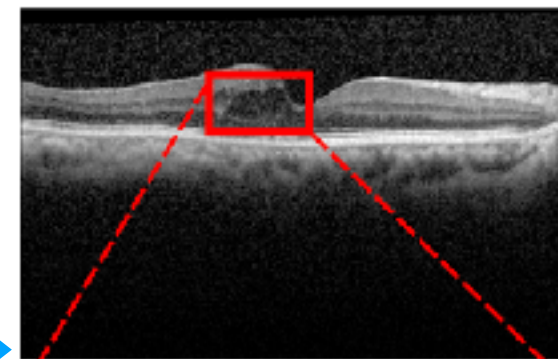
1st visit: **3D model** detects diabetes signal



1 year later



**2D model** detects diabetes

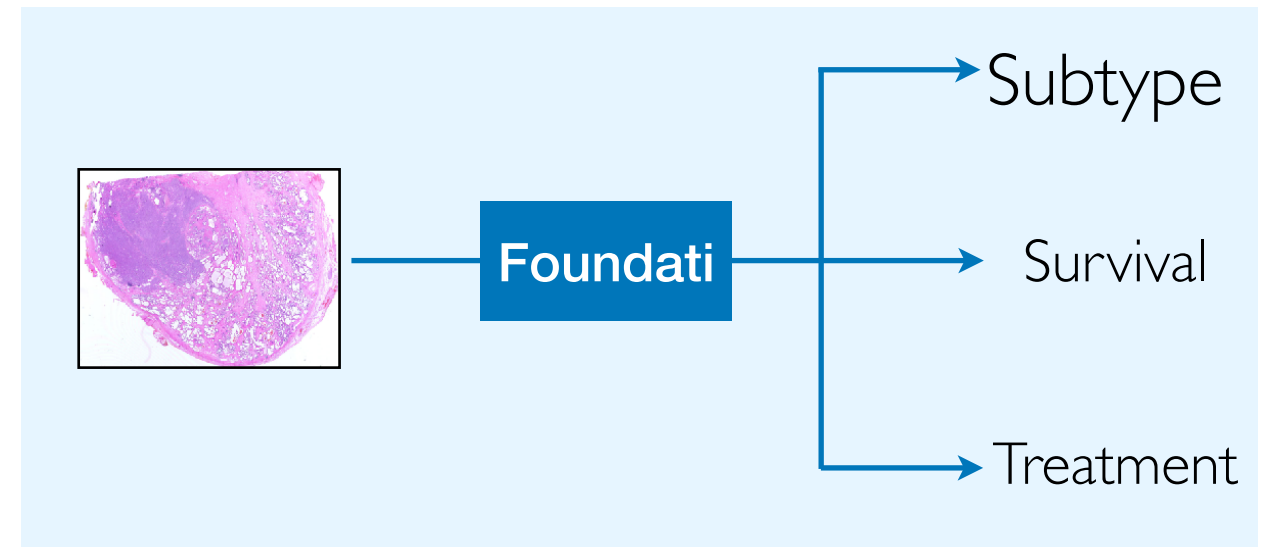


# Today's talk: 3 parts

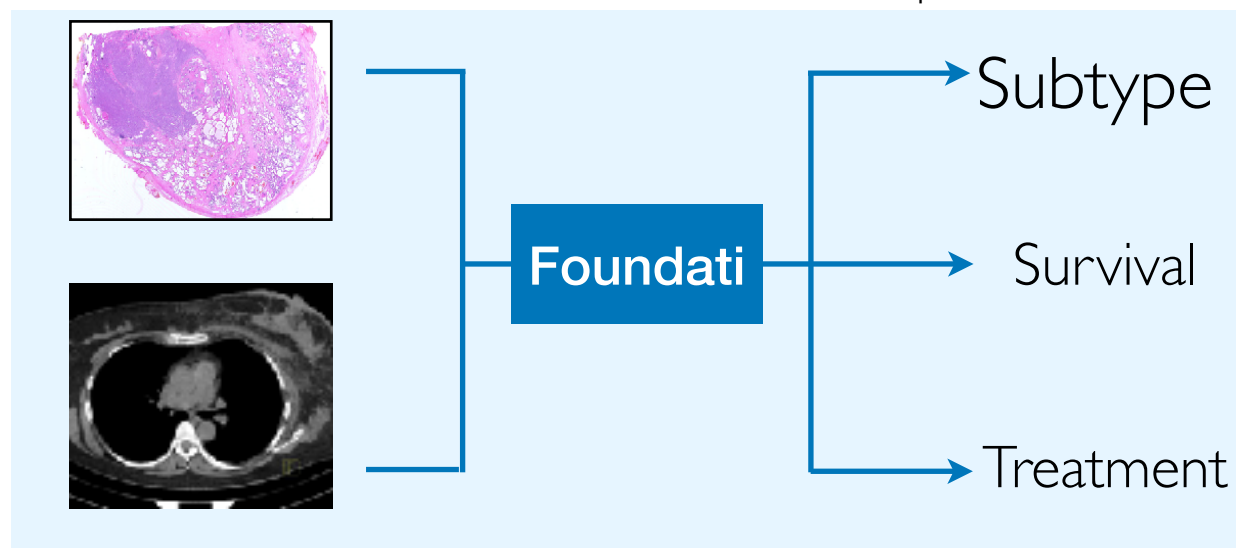
1. Pathology foundation model
2. 3D retinal foundation model
3. A multi-modal foundation model integrating 9 imaging modalities



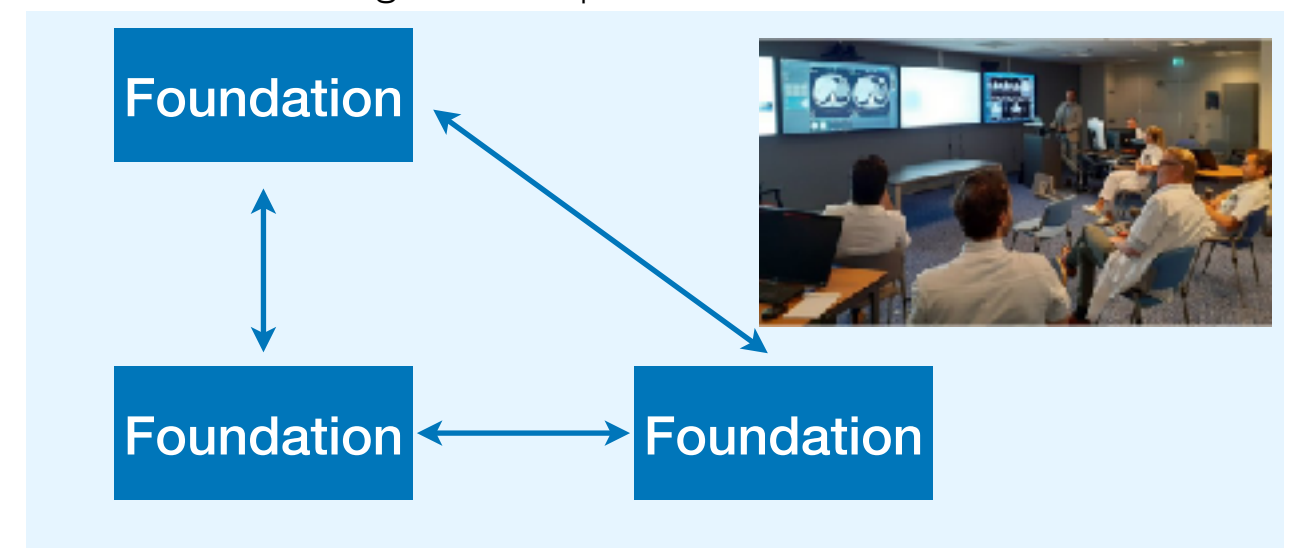
**Foundation model (2022)**  
One model for all tasks



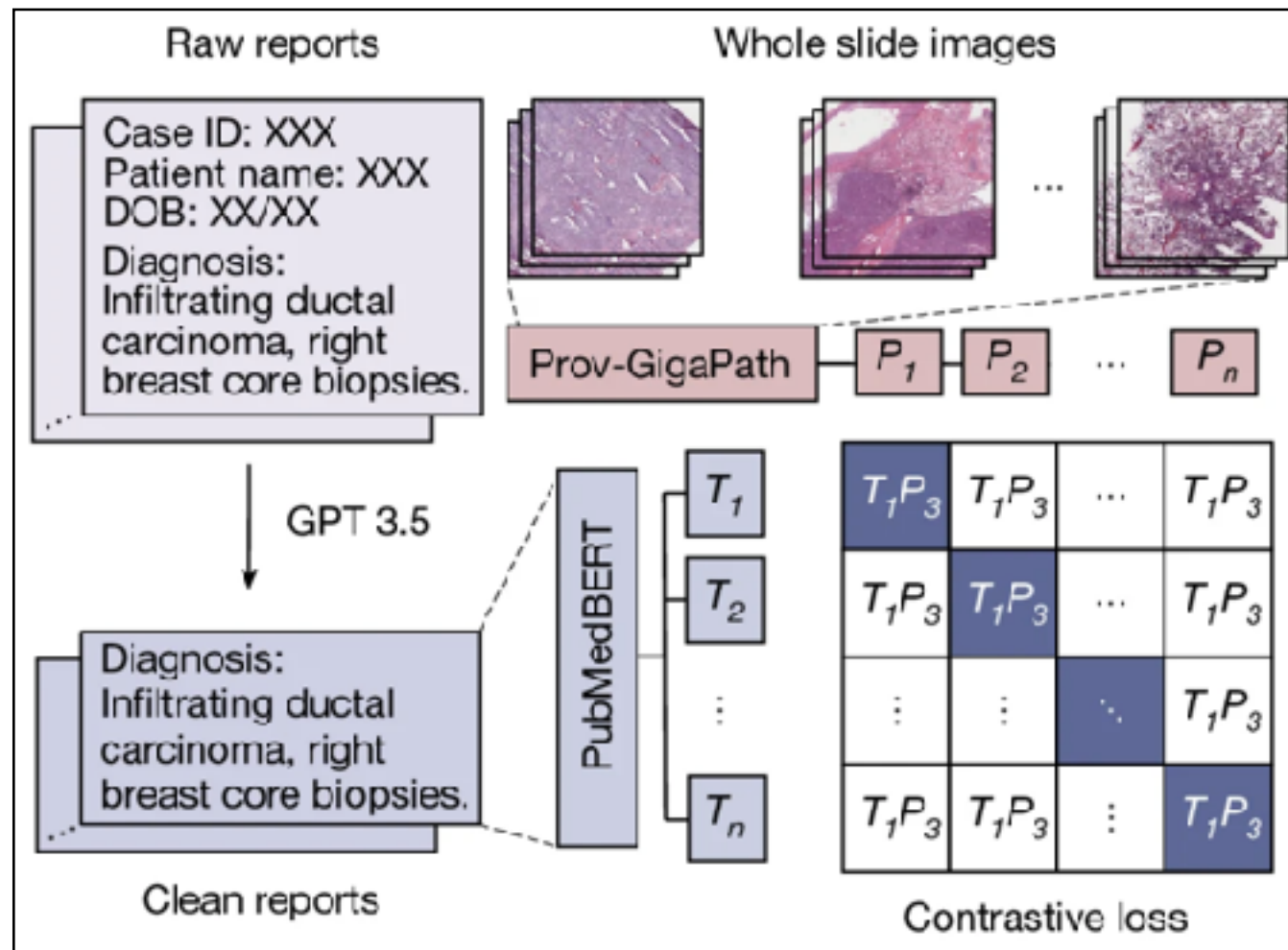
**Multi-modal foundation model (2023)**  
One model takes different inputs



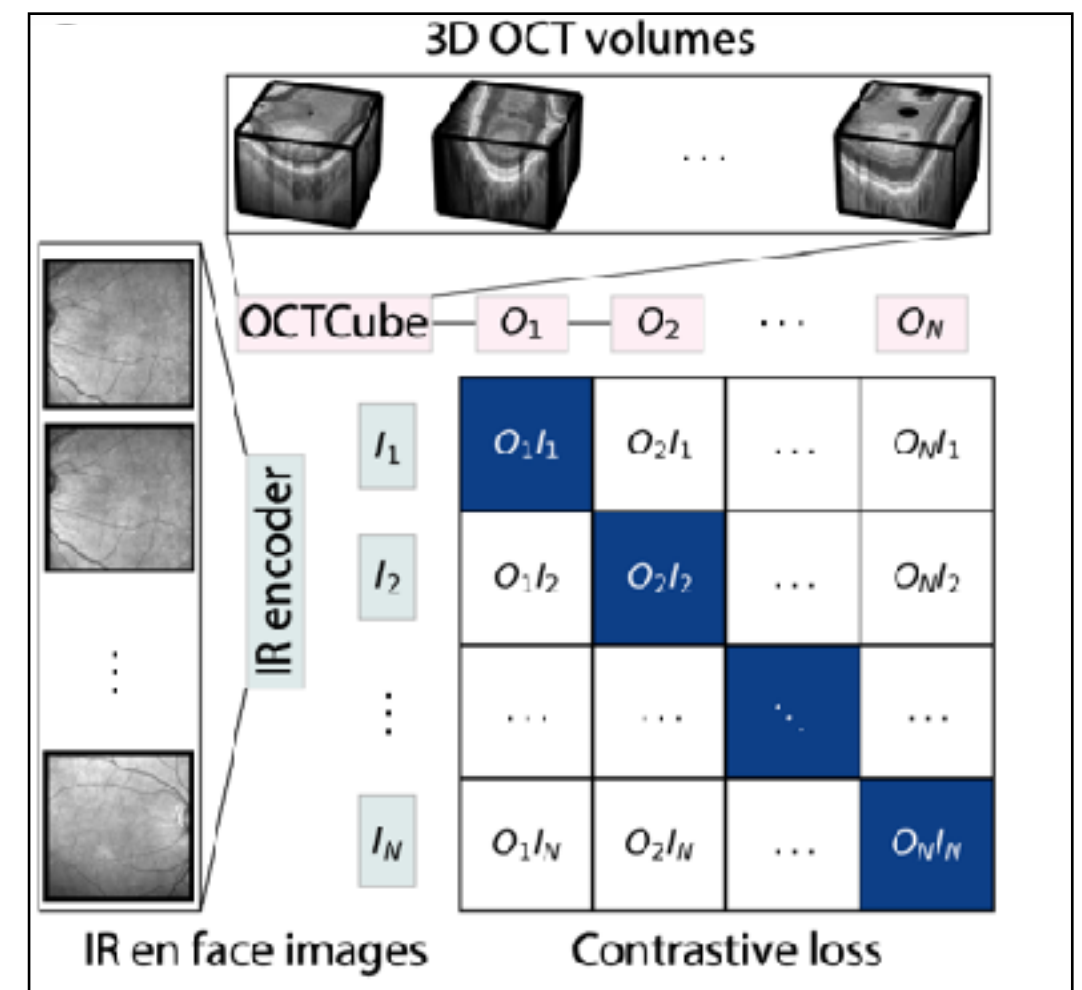
**Multi-agent (2024)**  
Integrate multiple foundation models



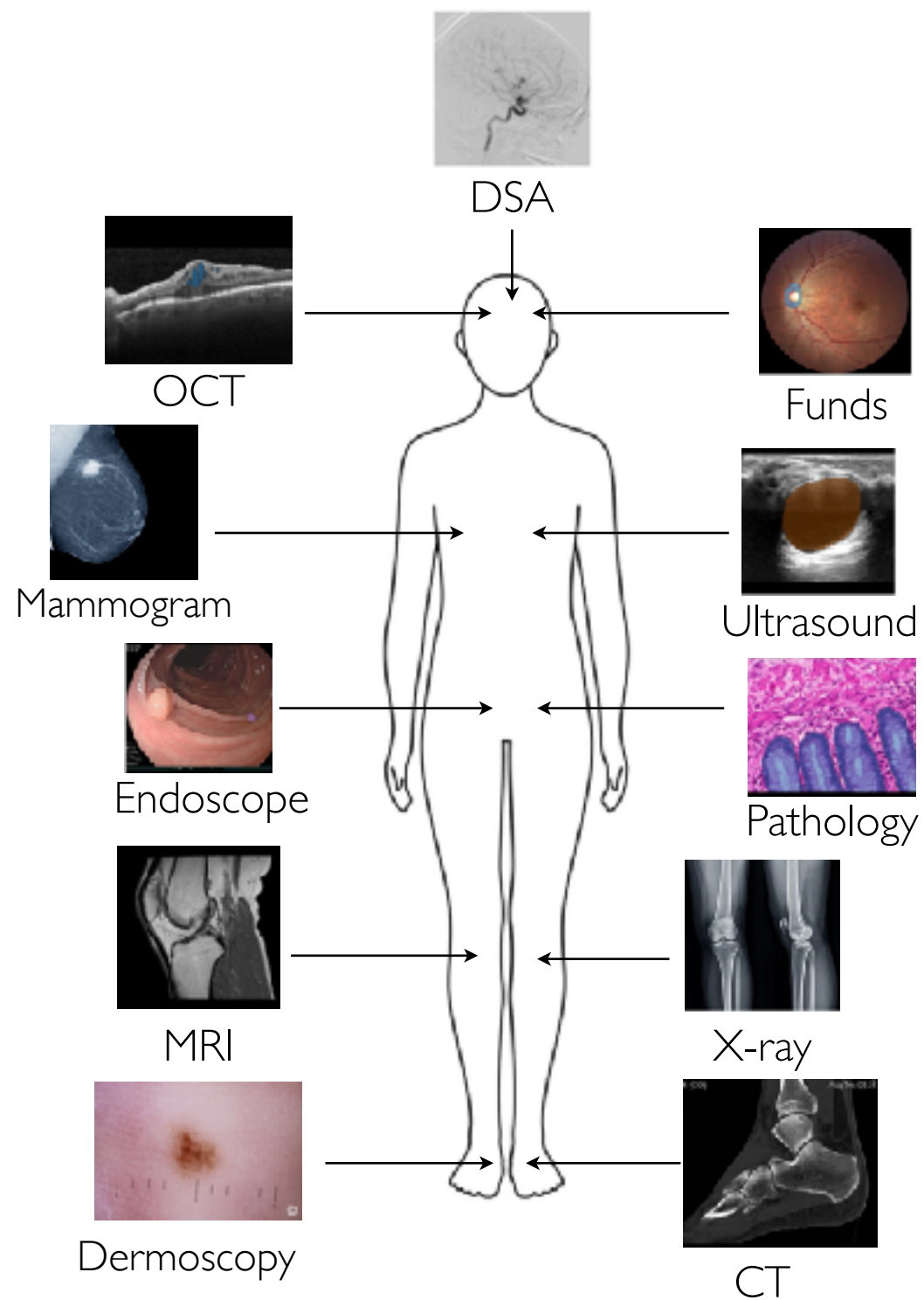
# Multi-modality analysis in GigaPath and OCTCube



Medical report + Pathology image



OCT image + IR image



Existing cross-modal framework is  
Limited to two modalities

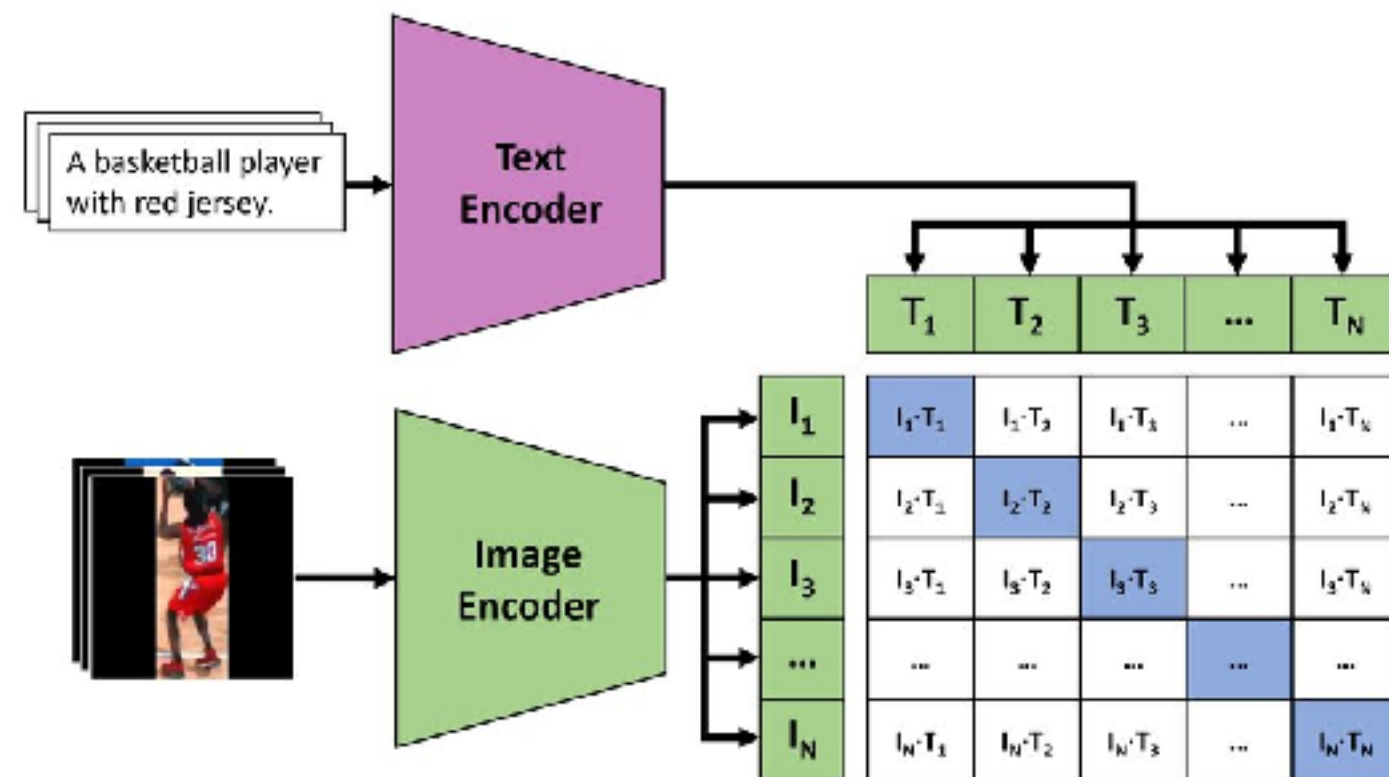
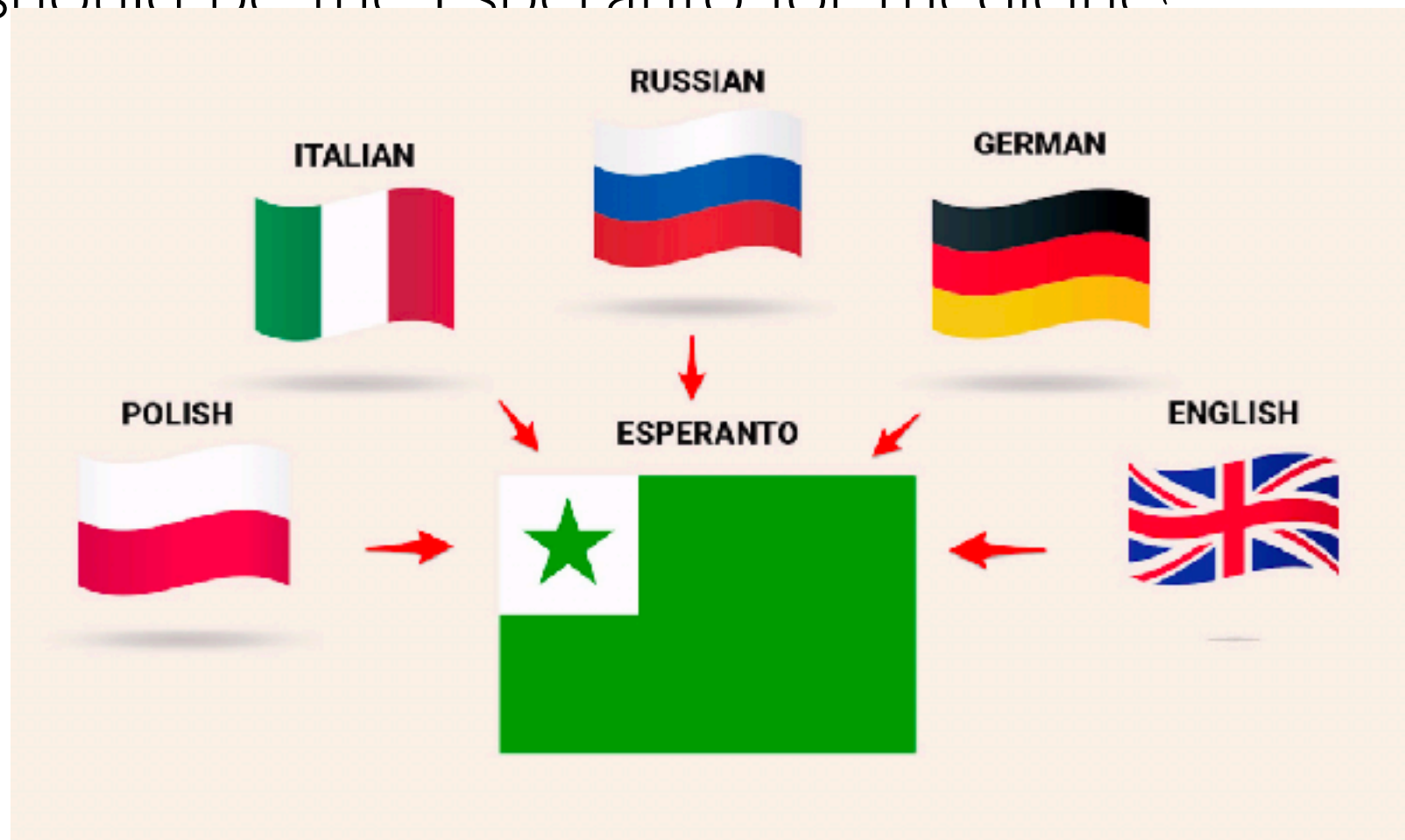


Image from CLIP-ReIdent

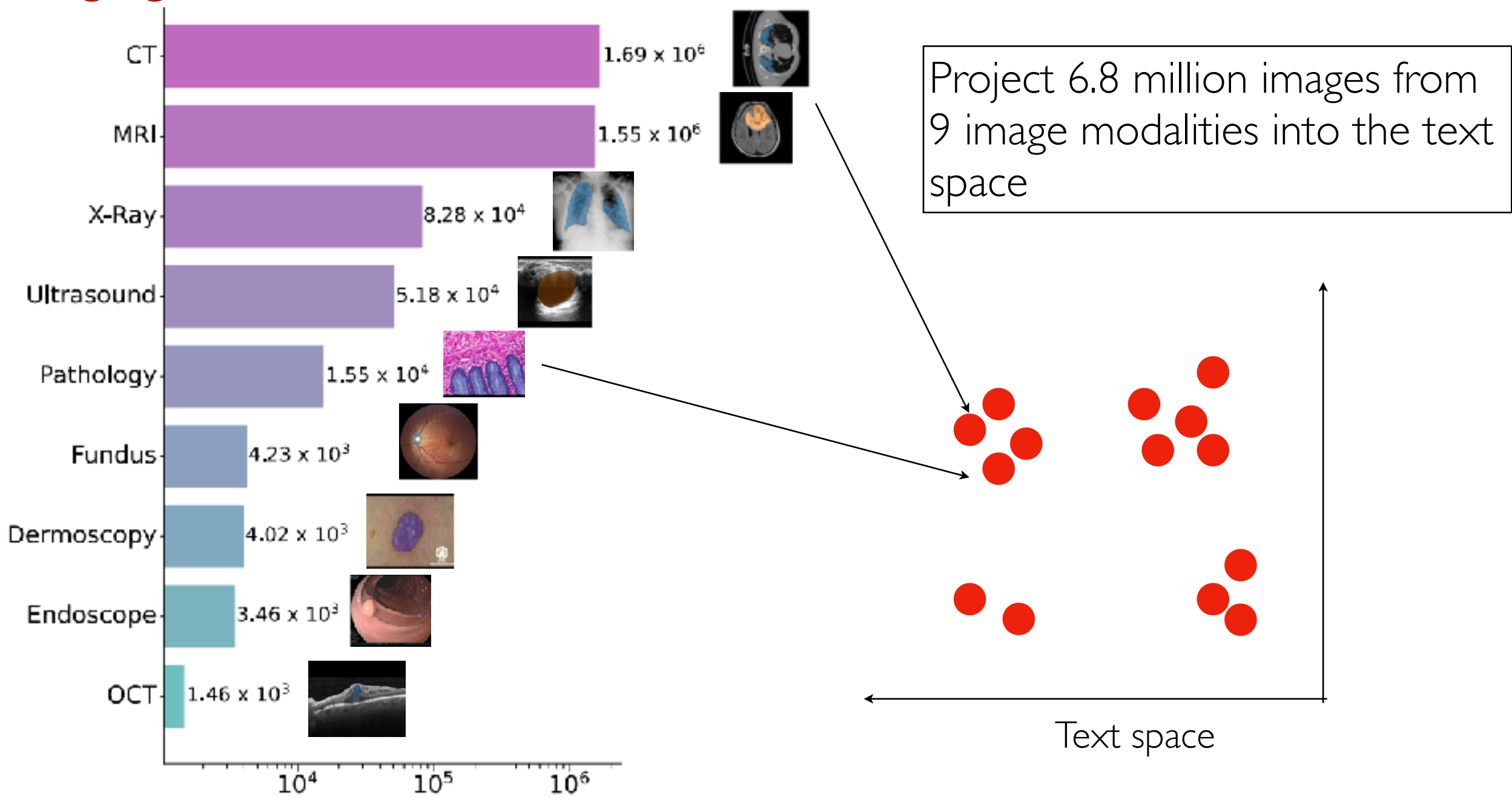
# Our solution for integrating all image modalities

- Build one foundation model for each modality, then project different models into the same space
- What should be the Esperanto for medicine?

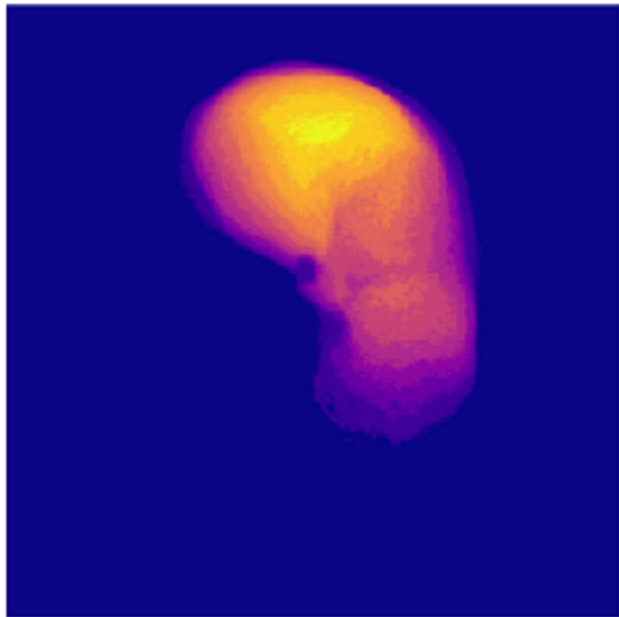




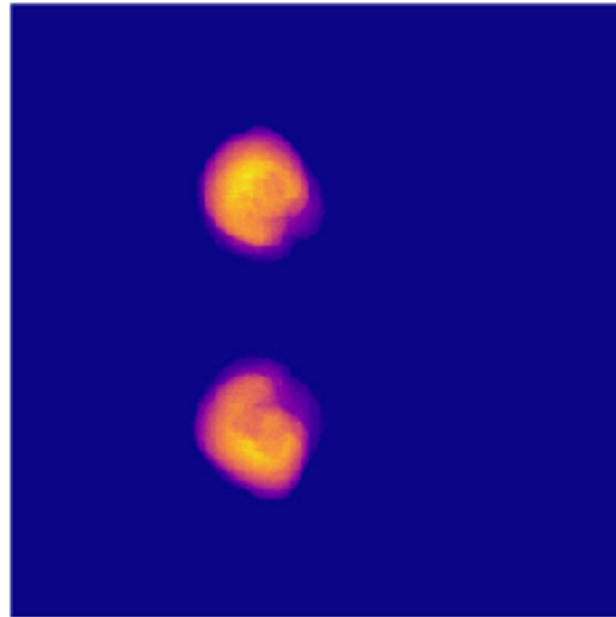
# BiomedParse: use human language as the Esperanto to integrate medical imaging modalities



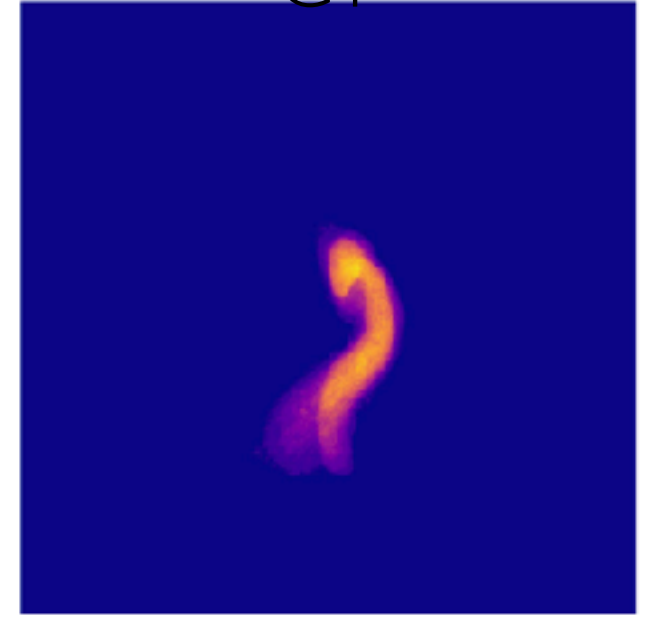
Liver in abdomen CT



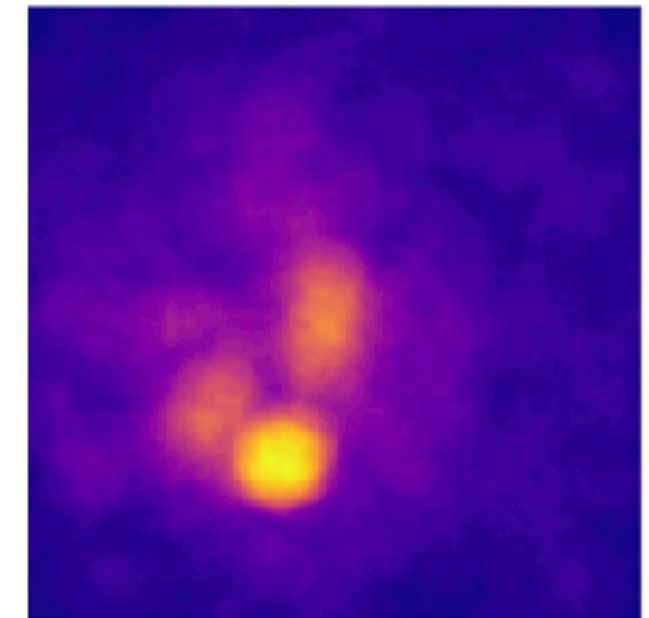
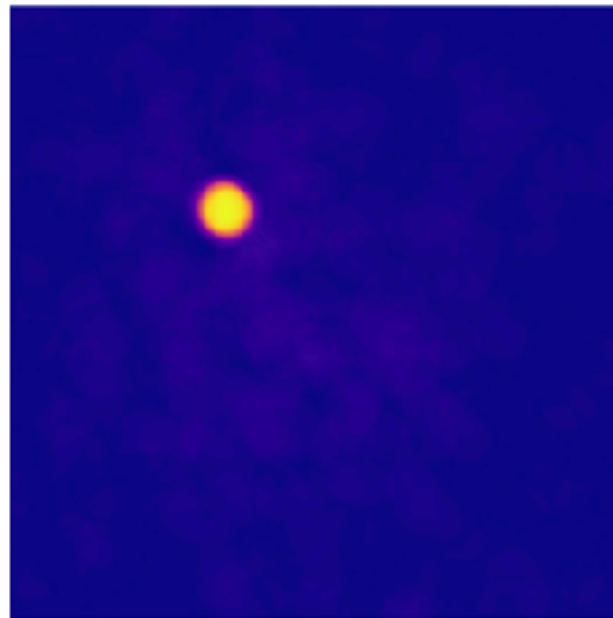
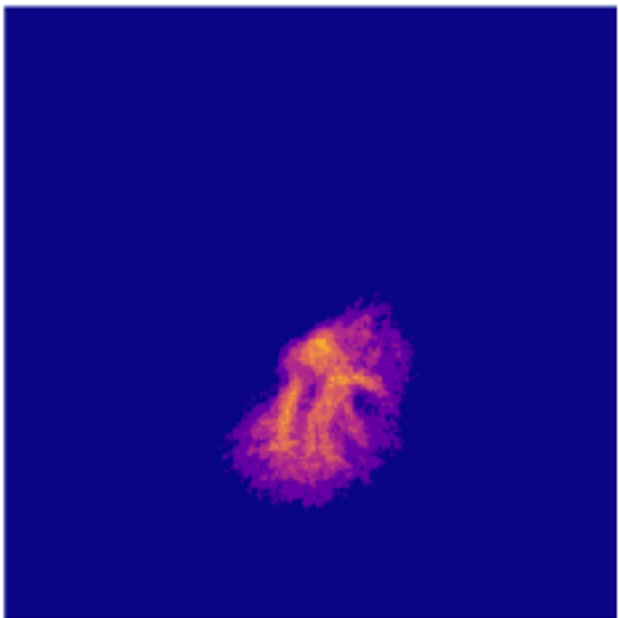
Kidney in abdomen CT



Pancreas in abdomen  
CT



Liver vessel in abdomen CT Inflammatory cells in pathology Neoplastic cells in pathology



# A foundation model for joint segmentation, detection and recognition across 9 modalities

nature methods

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Article | Published: 18 November 2024

## A foundation model for joint segmentation, detection and recognition of biomedical objects across nine modalities

[Theodore Zhao](#), [Yu Gu](#), [Jianwei Yang](#), [Naoto Usuyama](#), [Ho Hin Lee](#), [Sid Kiblawi](#), [Tristan Naumann](#),  
[Jianfeng Gao](#), [Angela Crabtree](#), [Jacob Abel](#), [Christine Mounq-Wen](#), [Brian Piening](#), [Carlo Bifulco](#), [Mu Wei](#)  
✉, [Hoifung Poon](#) ✉ & [Sheng Wang](#) ✉

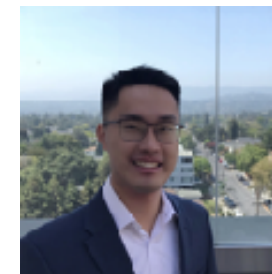
[Nature Methods](#) (2024) | [Cite this article](#)



Theodore Zhao



Aiden Gu



Sheng Wang  
U of Washington



Mu Wei

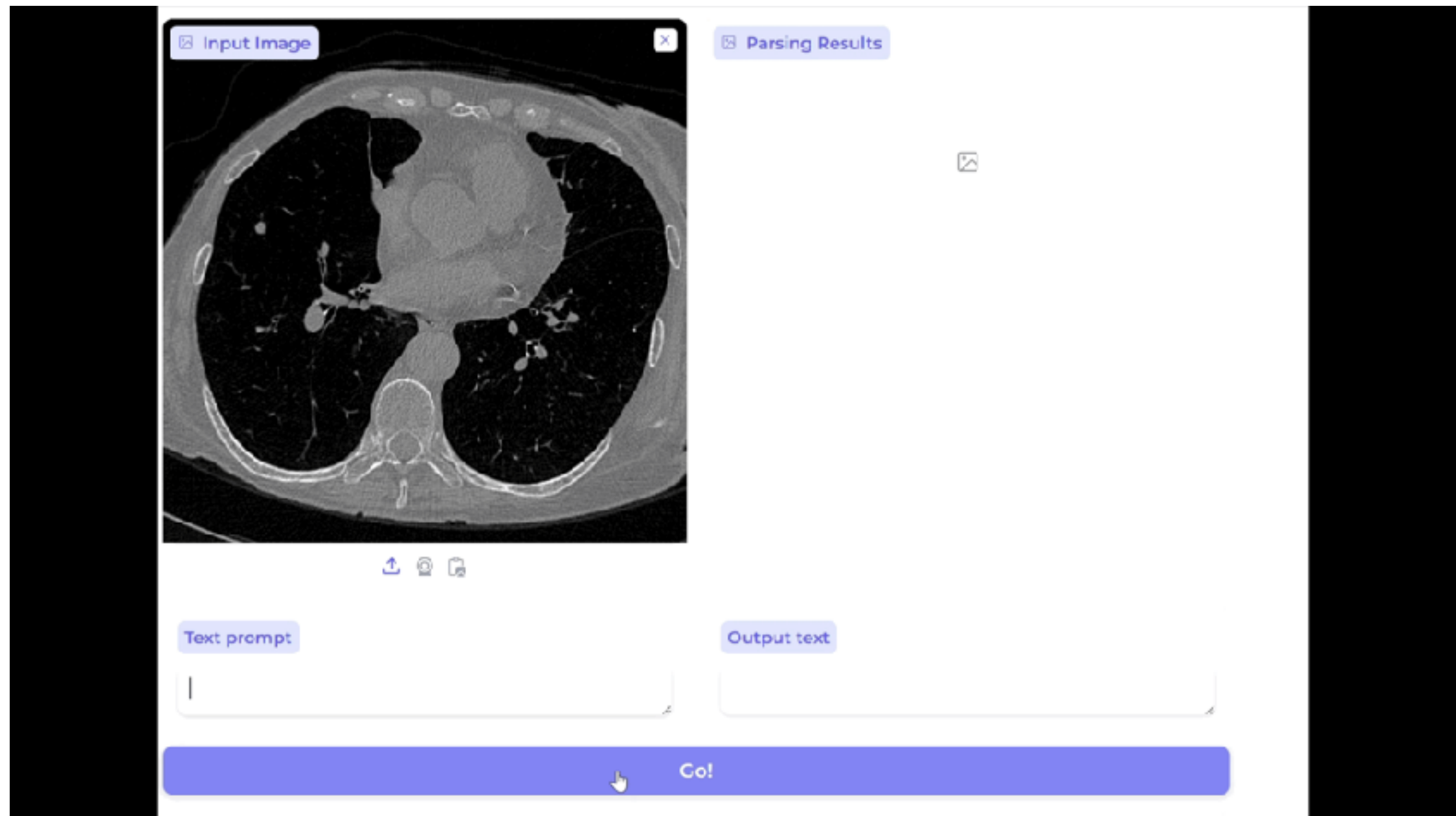


Hoifung Poon

Available as open-source model on Azure AI and Hugging Face



## Demo I: talk to the AI model to find lung nodule

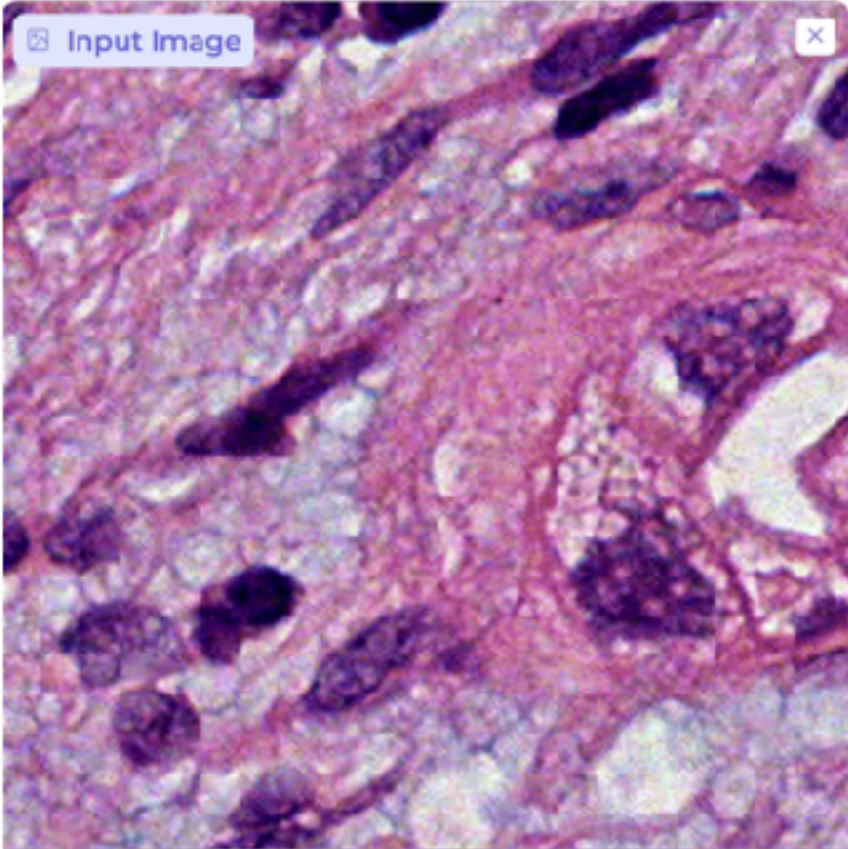





## Demo 1: talk to the AI model to find lung nodule



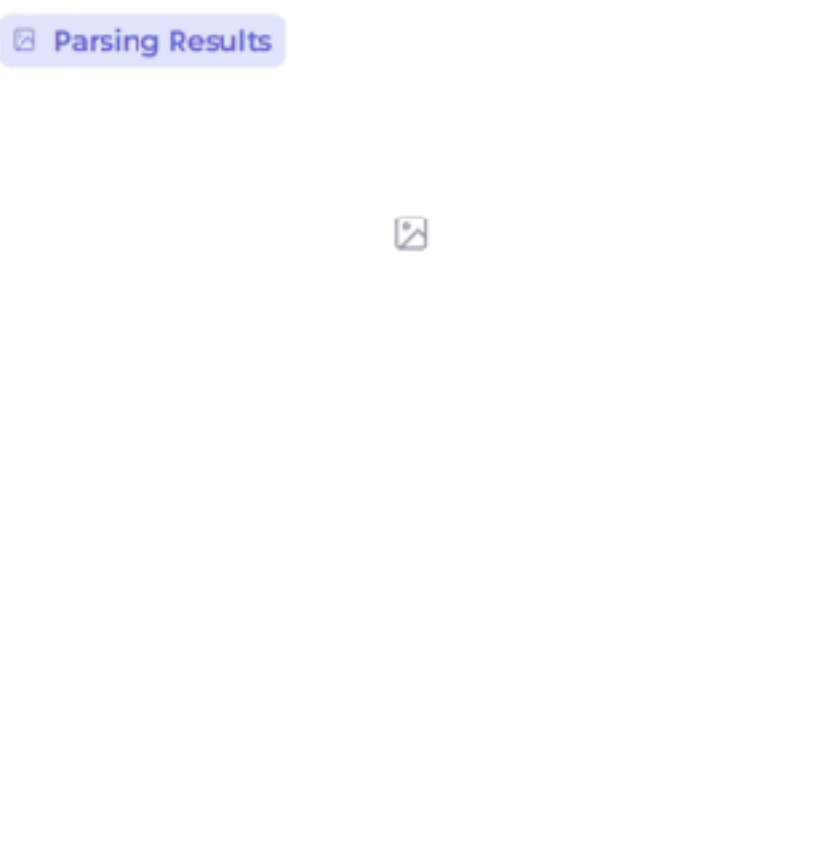
## Demo2: talk to the model to find all cell types

Input Image





Parsing Results



Text prompt

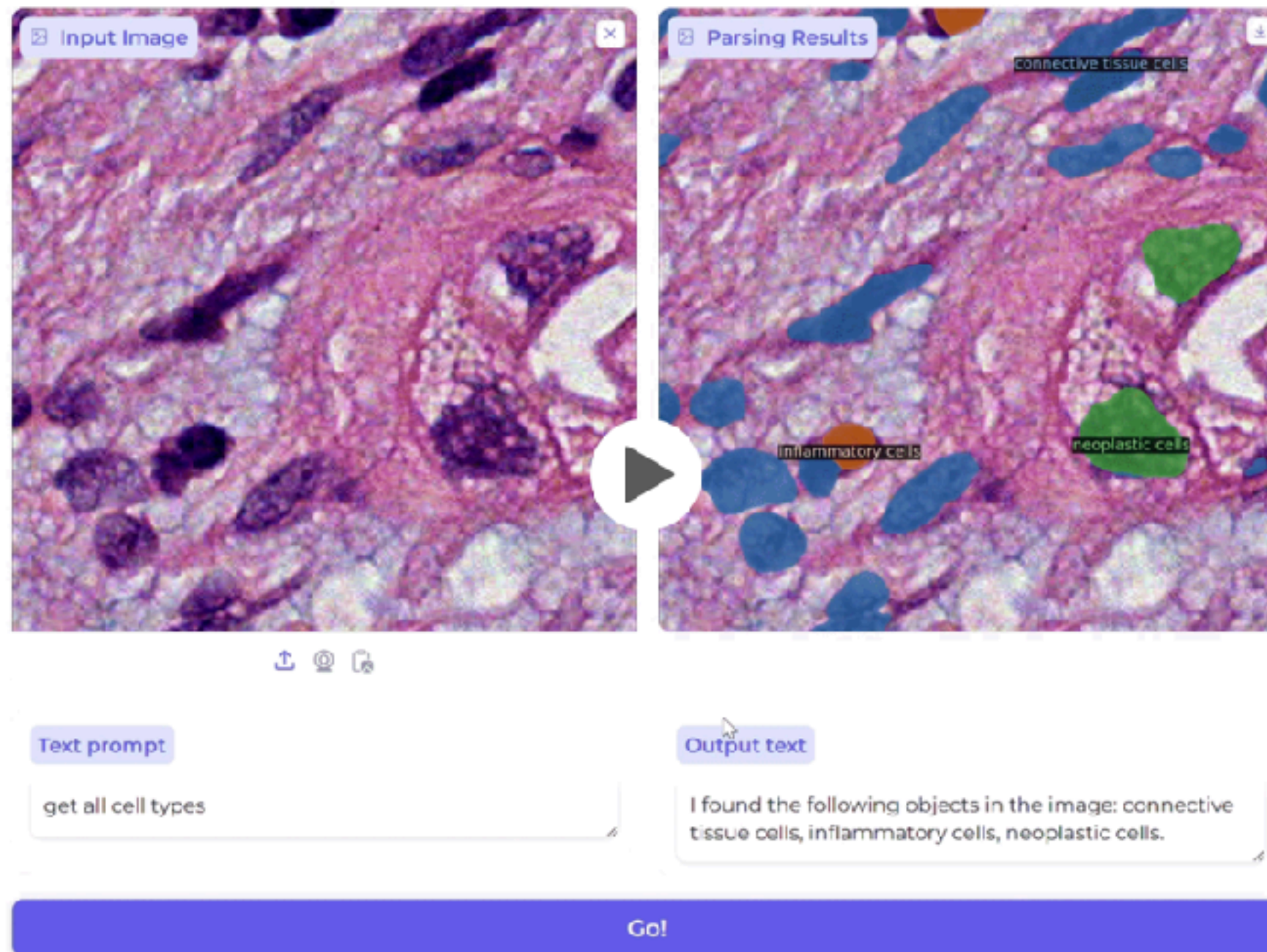
g

Output text

Go!



## Demo2: talk to the model to find all cell types



The interface is divided into two main columns. The left column, titled "Input Image", shows a histological slide. The right column, titled "Parsing Results", shows the same slide with colored overlays and labels: "connective tissue cells" (blue), "inflammatory cells" (orange), and "neoplastic cells" (green). A play button is centered between the two images. Below the images are two text input fields. The left field, labeled "Text prompt", contains the text "get all cell types". The right field, labeled "Output text", contains the text "I found the following objects in the image: connective tissue cells, inflammatory cells, neoplastic cells." A blue "Go!" button is at the bottom.

Input Image

Parsing Results

connective tissue cells

inflammatory cells

neoplastic cells

Text prompt

get all cell types

Output text

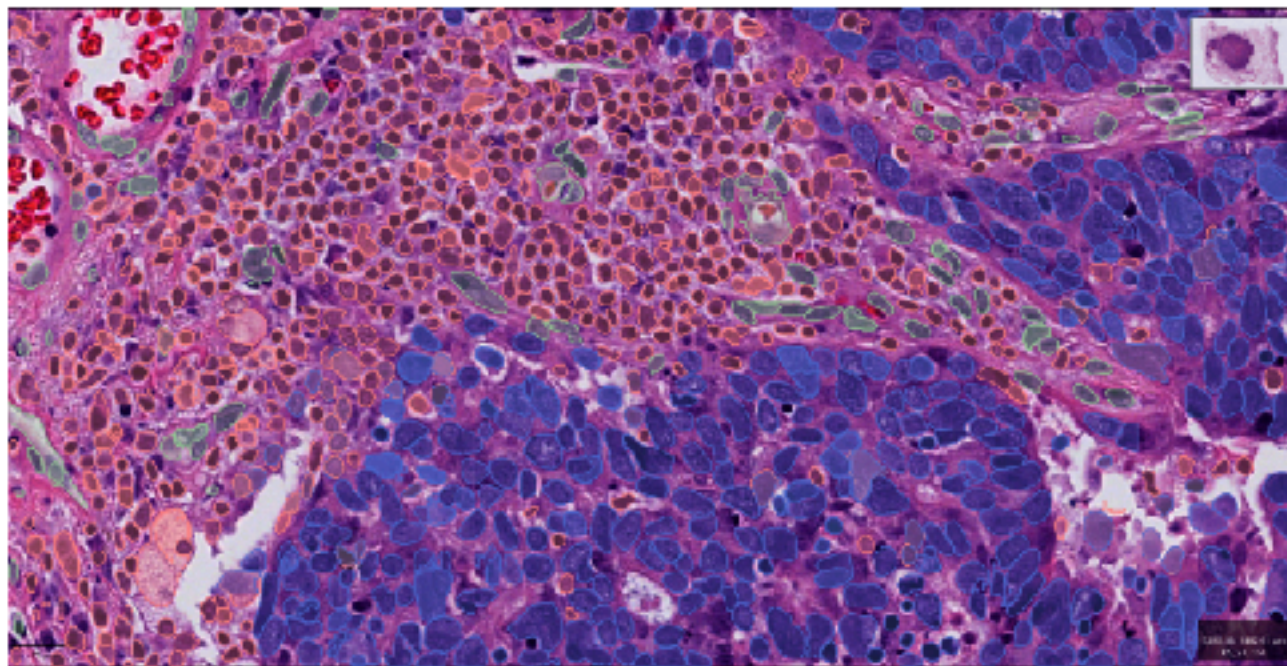
I found the following objects in the image: connective tissue cells, inflammatory cells, neoplastic cells.

Go!



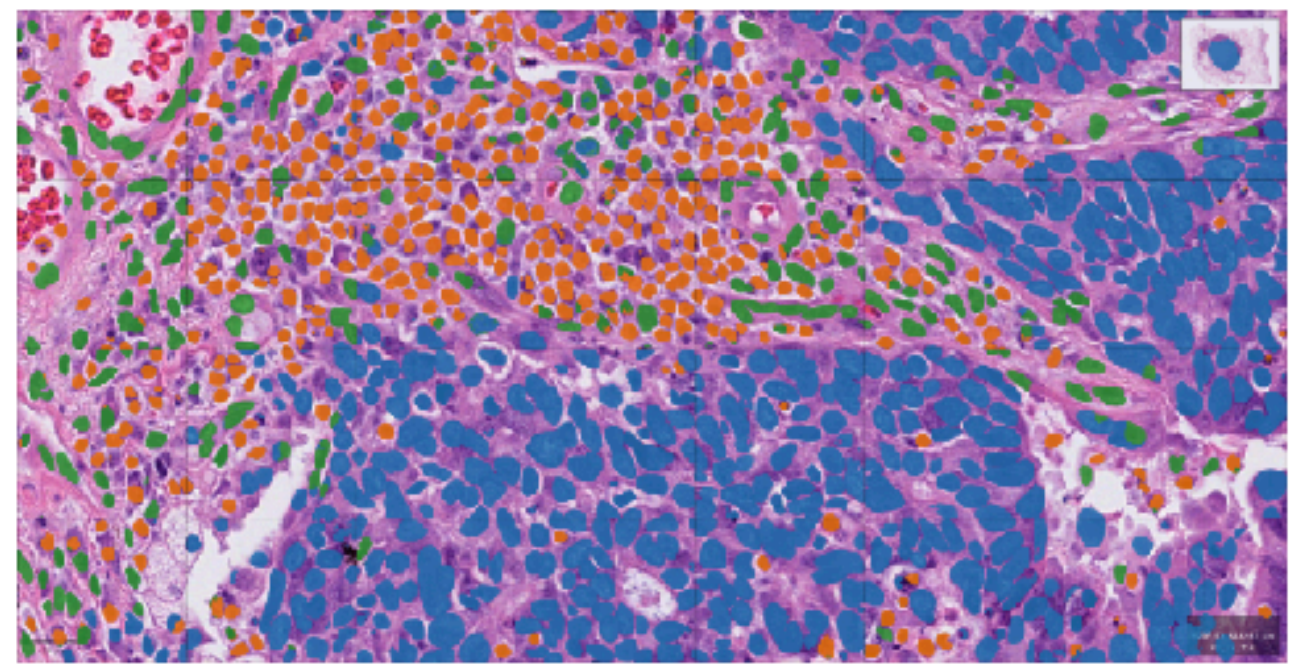
0.2 second to segment and label all cells with >  
90% consistency

Pathologist annotation



 Tumor region  Lymphocyte  Stroma

BiomedParse prediction

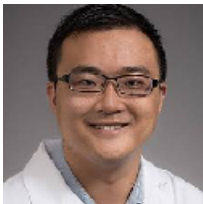
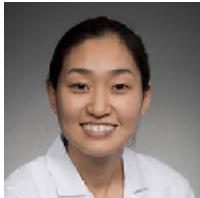
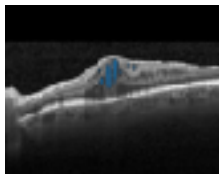


 Neoplastic cells  Inflammatory cells  
 Connective tissue cells



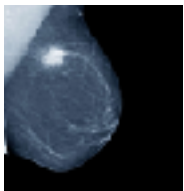
# Data-raising from Collaborators at UW School of Medicine

27k OCT images for retinal diseases



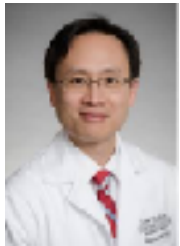
Drs. Cecilia and Aaron Lee (Ophthalmology)

50k Mammogram for breast cancer



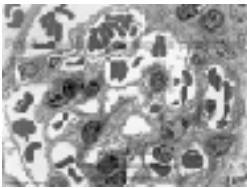
Dr. Christopher Lee (Radiology)

100k CT for heart transplant



Dr. Shin Lin (Cardiology)

190k EM image for kidney



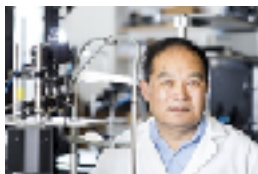
Dr. Behzad Najafian (Pathology)



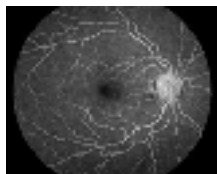
Dr. Mehmet Kurt (ME)



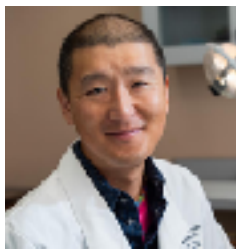
100k brain MRI for stroke, Parkinson, brain cancer



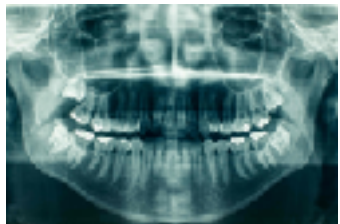
Dr. Ricky Wang (BioE)



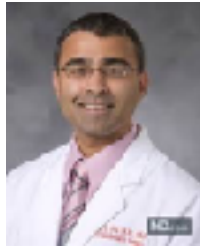
5k FA for glaucoma



Drs. Donald Chi, Amy Kim (Dentistry)



50k dental panoramic X-ray



Dr. Jay Pal (Surgery)



300k echo for heart failure



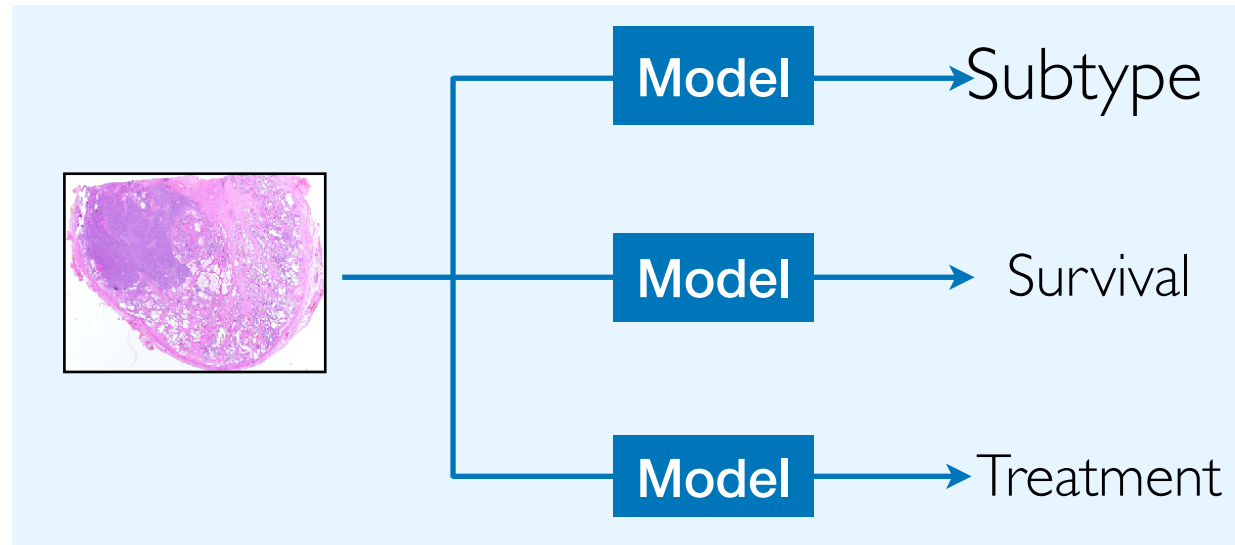
Drs. Nathan Cross and Paul Kinahan (Radiology)



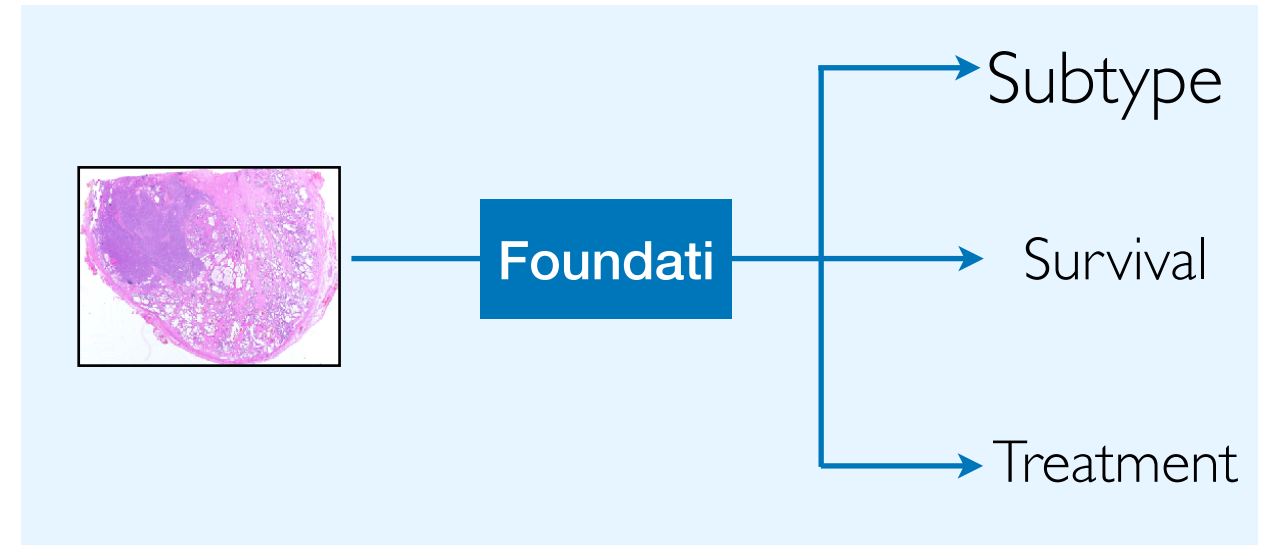
120k spinal MRI for back pain

# Four paradigms in AI for Medicine

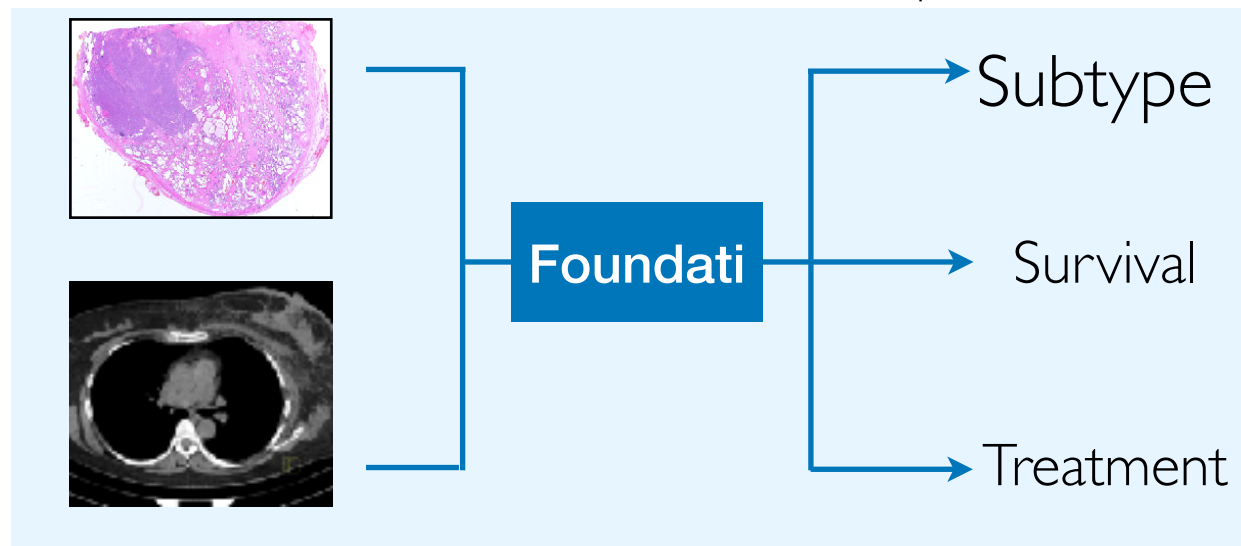
Deep learning (2012)  
One model for one task



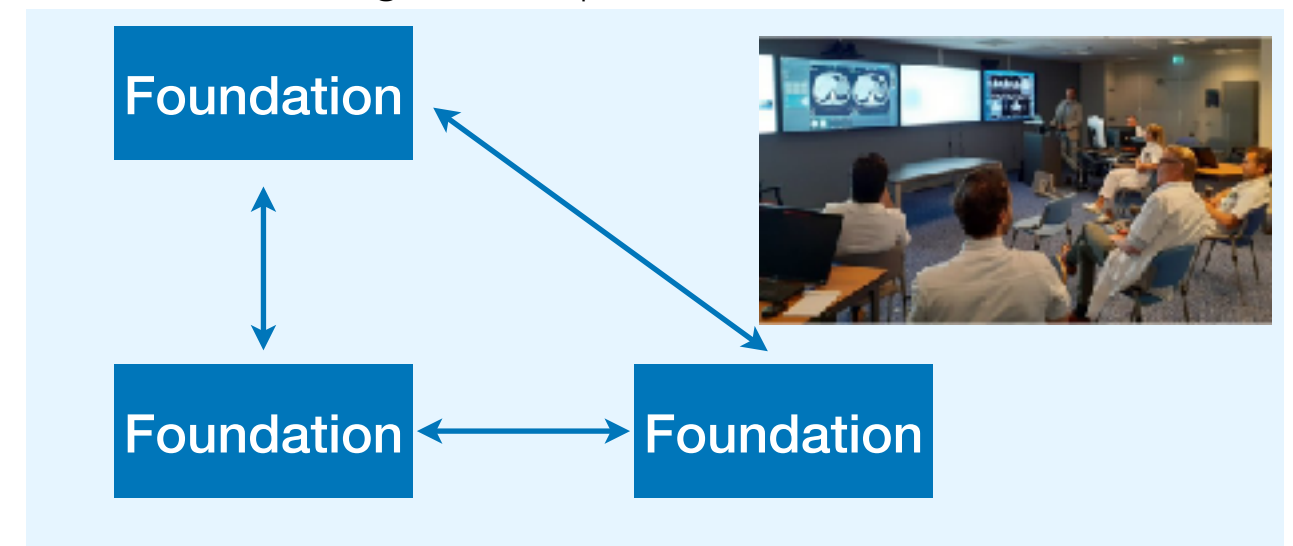
Foundation model (2022)  
One model for all tasks



Multi-modal foundation model (2023)  
One model takes different inputs

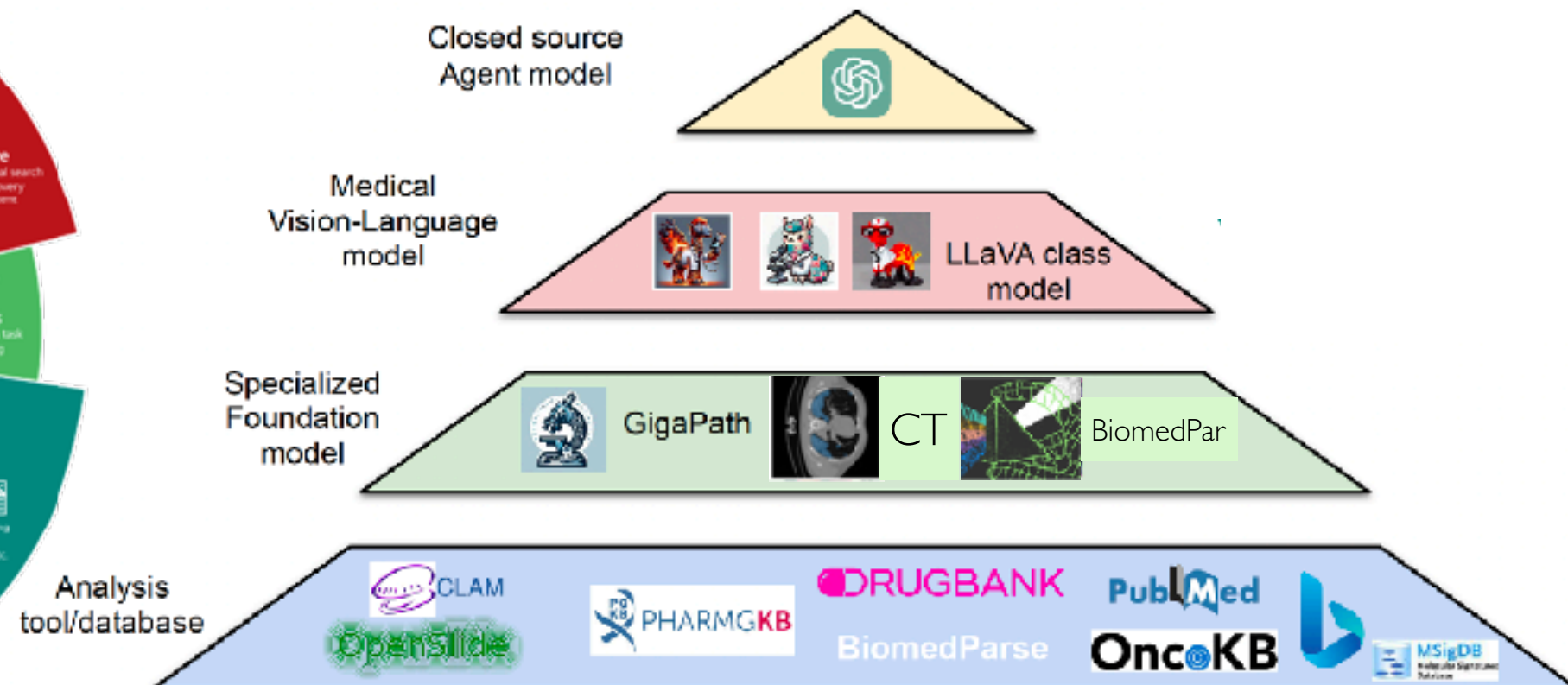


Multi-agent (2024)  
Integrate multiple foundation models





# A Microsoft 365 for cancer diagnosis



## Molecular tumor board: From multi-modality to multi-agent

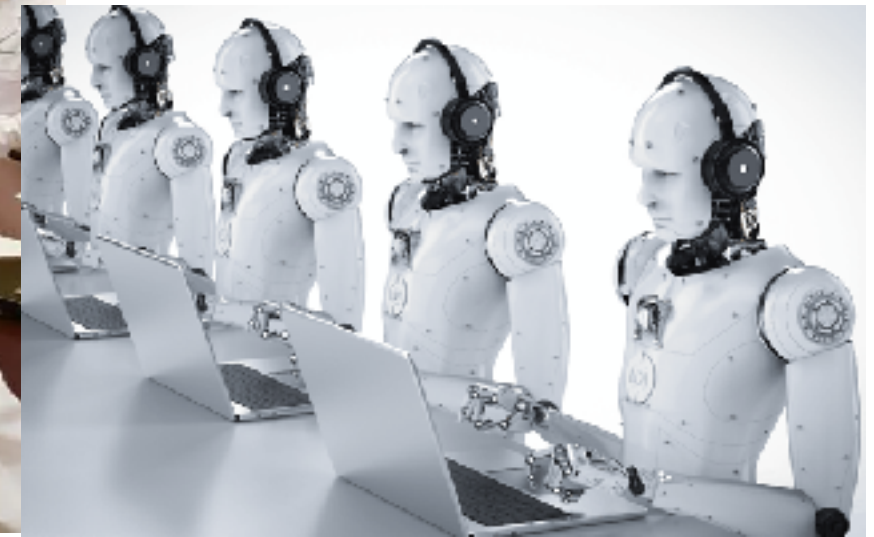
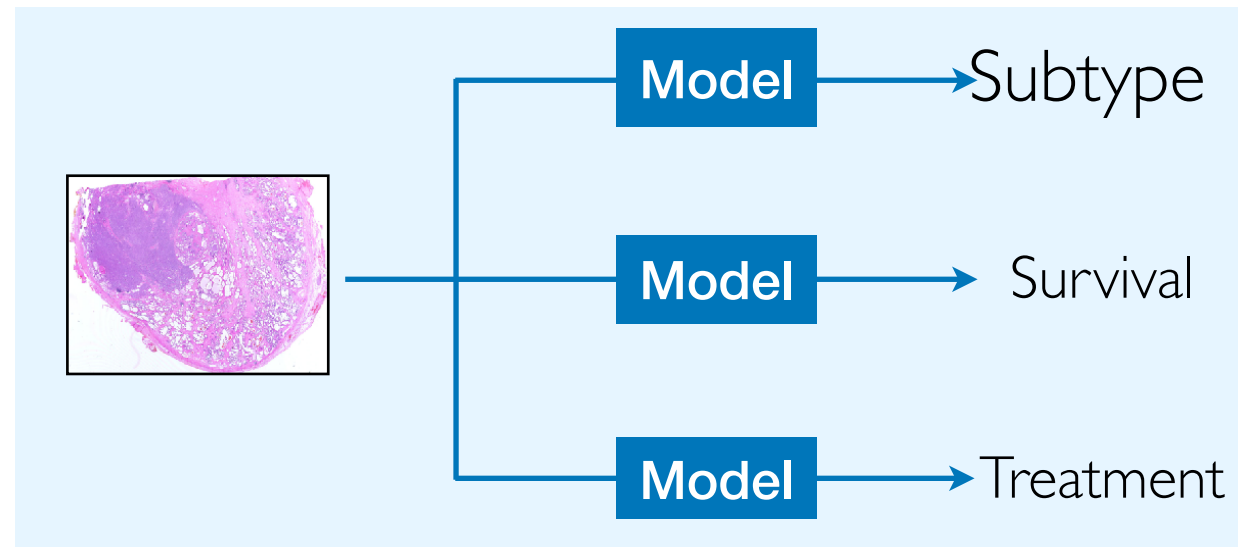


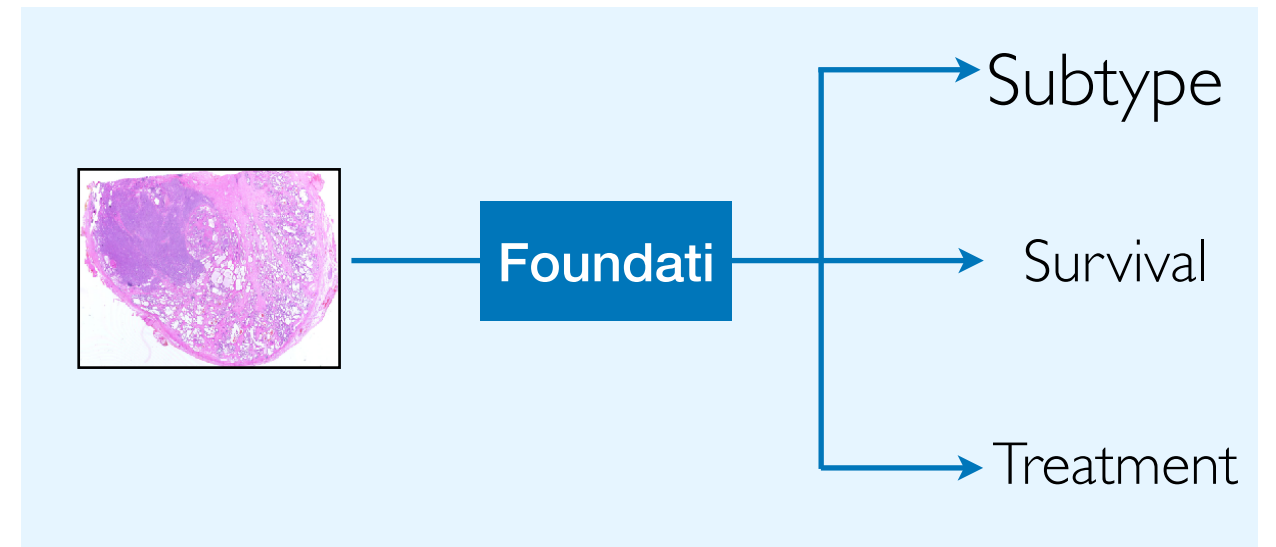
Image source: Mercy Health Brain Tumor Center

# Four paradigms in AI for Medicine

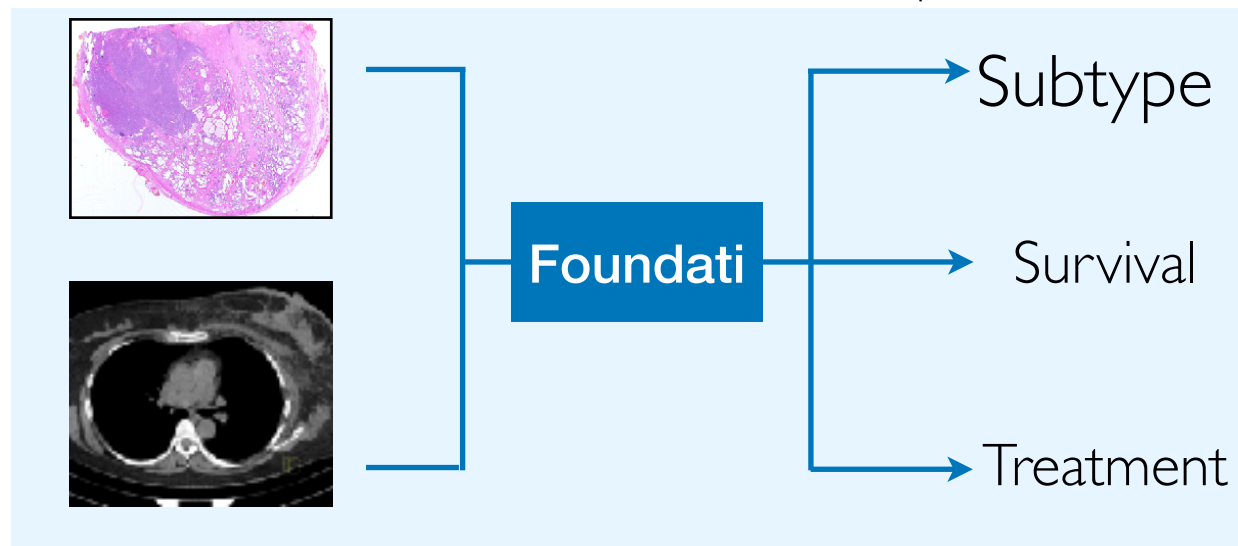
Deep learning (2012)  
One model for one task



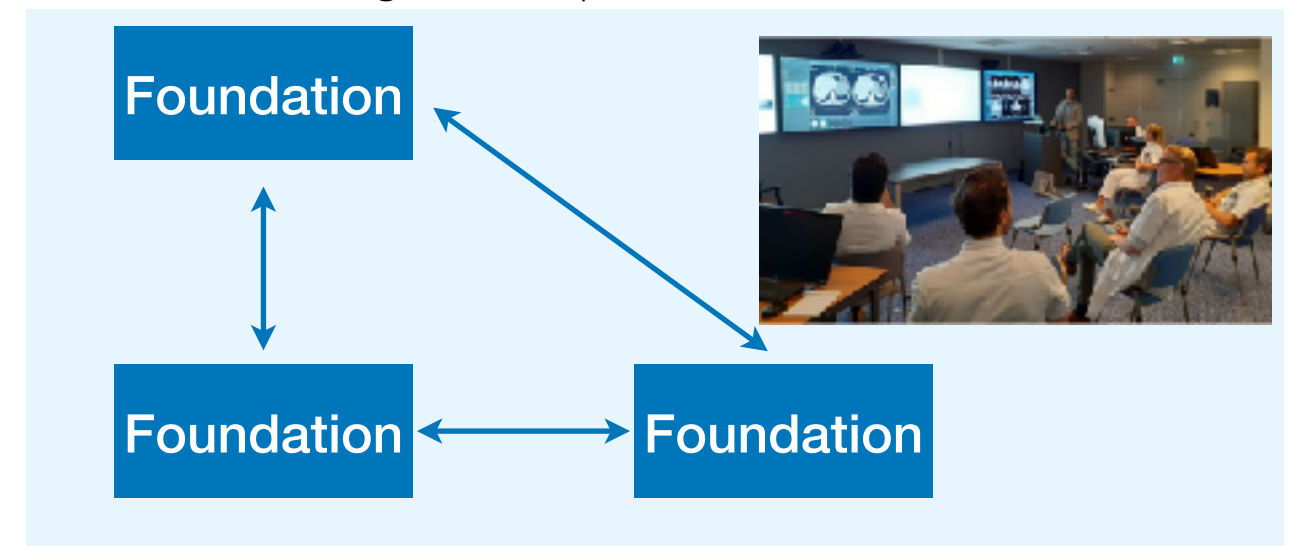
Foundation model (2022)  
One model for all tasks



Multi-modal foundation model (2023)  
One model takes different inputs



Multi-agent (2024)  
Integrate multiple foundation models



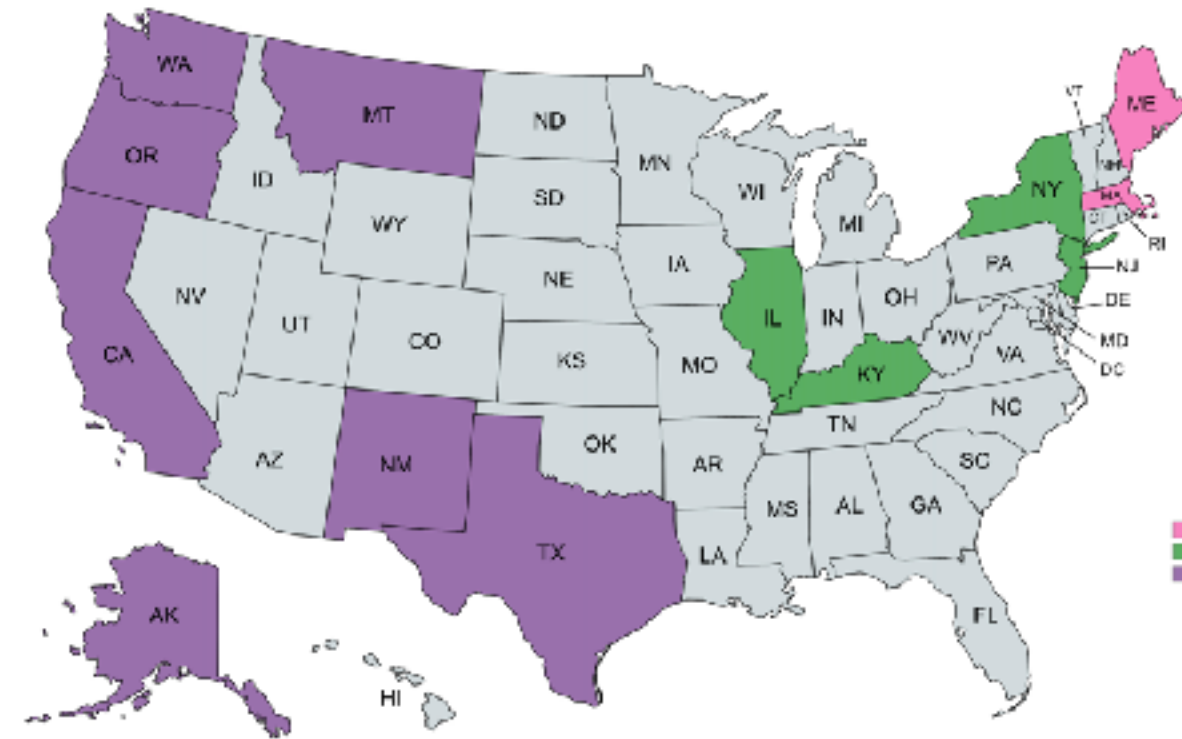
## Three lessons we learnt from GenAI for Medicine

- ✓ Medical foundation models are accurate

## Three lessons we learnt from GenAI for Medicine

- ✓ Medical foundation models are accurate
- ✓ Medical foundation models are heterogenous

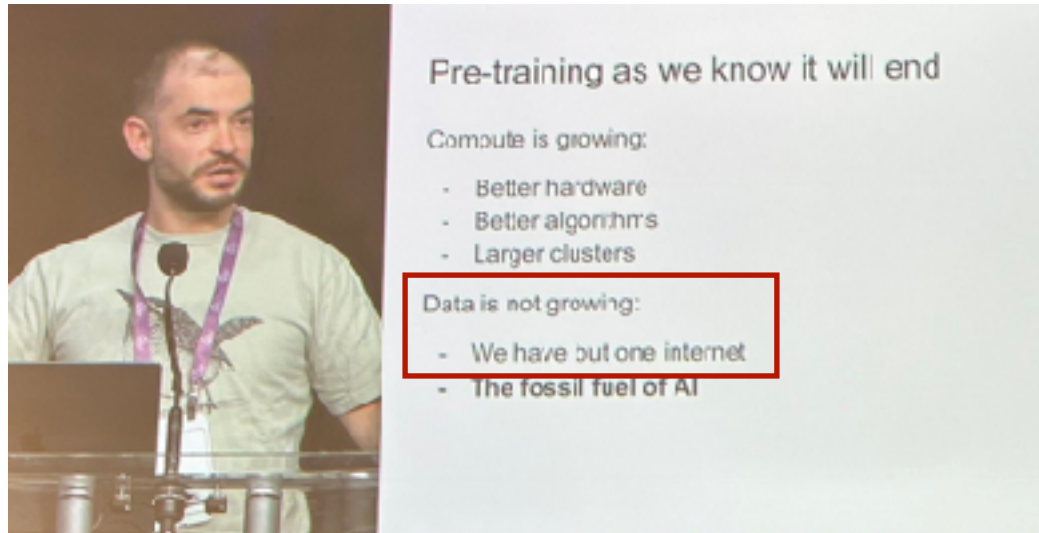
Data sources of three existing pathology models



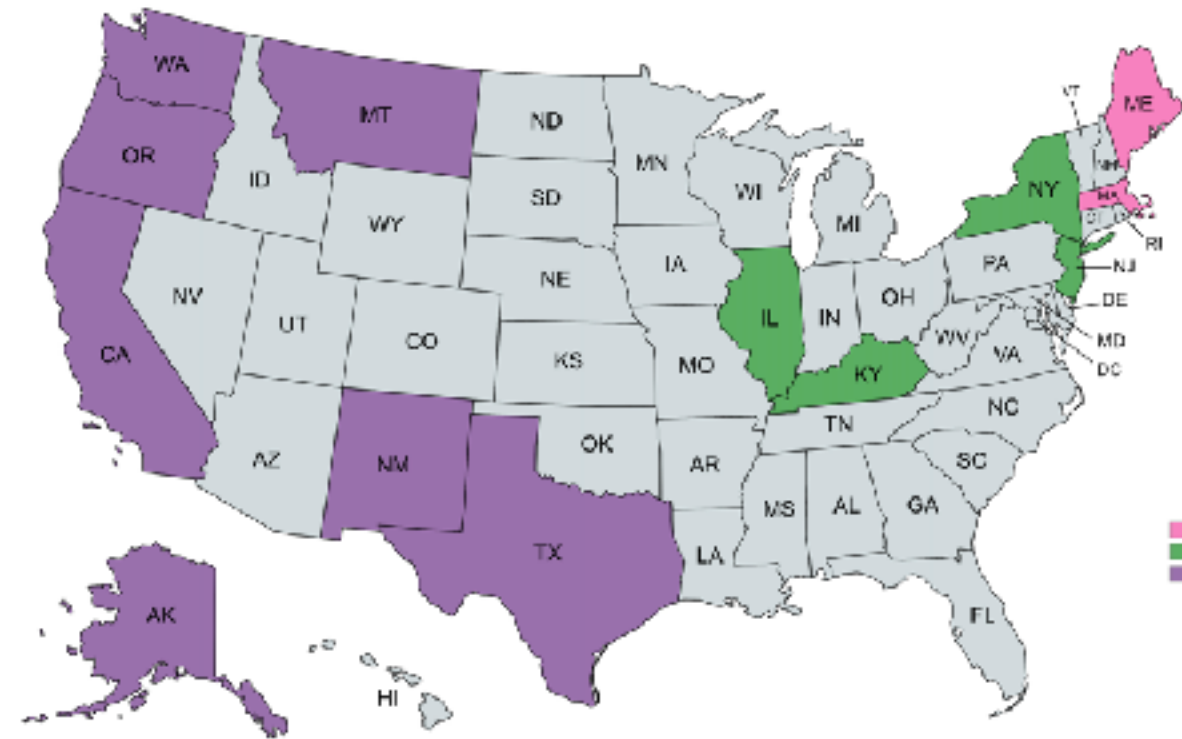


## Three lessons we learnt from GenAI for Medicine

- ✓ Medical foundation models are accurate
- ✓ Medical foundation models are heterogenous



Data sources of three existing pathology models

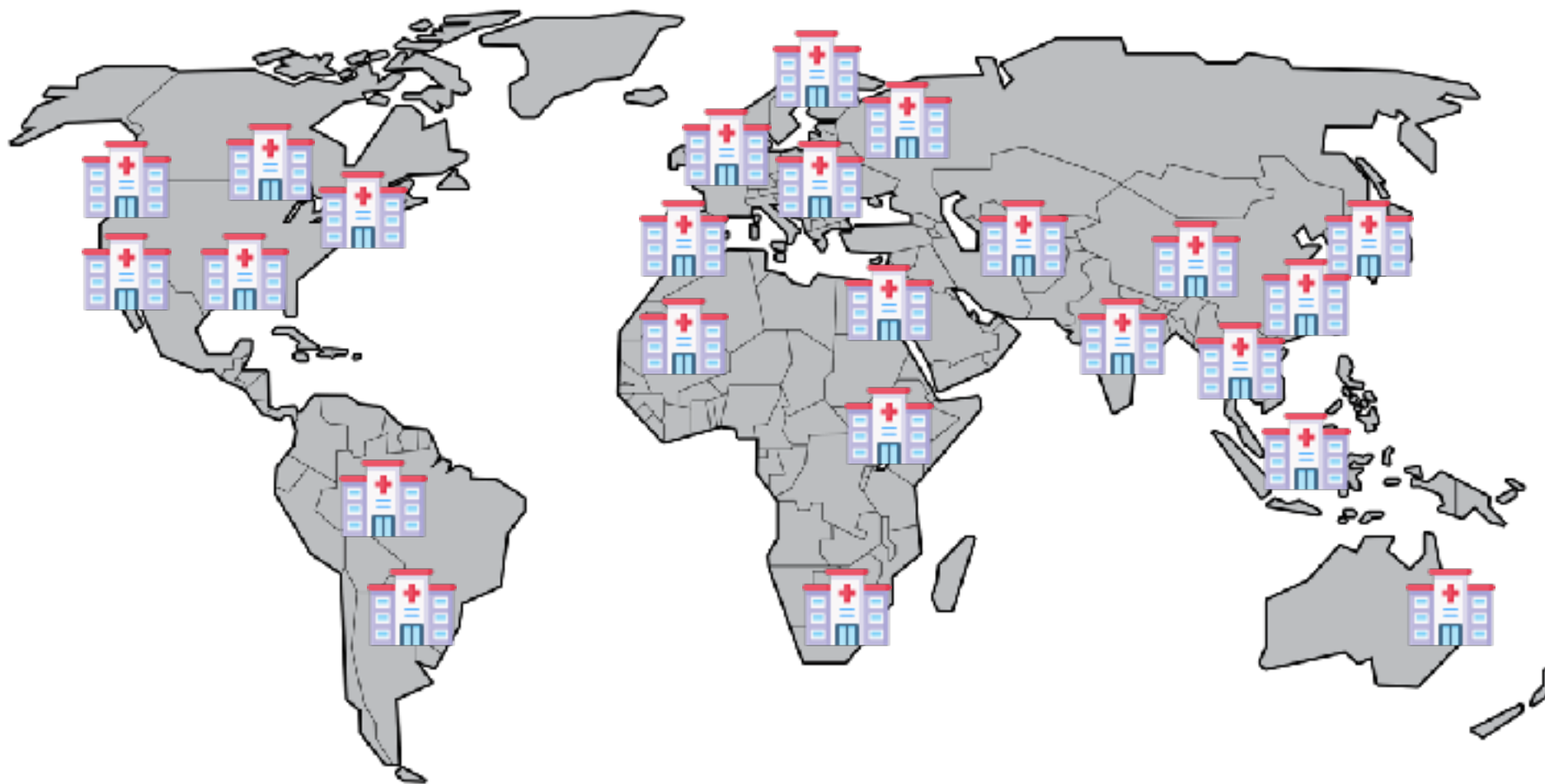


- ✓ **Less than 0.001% existing data** is used by even the largest medical foundation model
  - CT: 80m per year in US **vs.** 100k (10 years) by largest CT model
  - Pathology: 100m slides per year in US **vs.** 170k slides (10 years) by largest pathology model



## Fifth paradigm: a world model

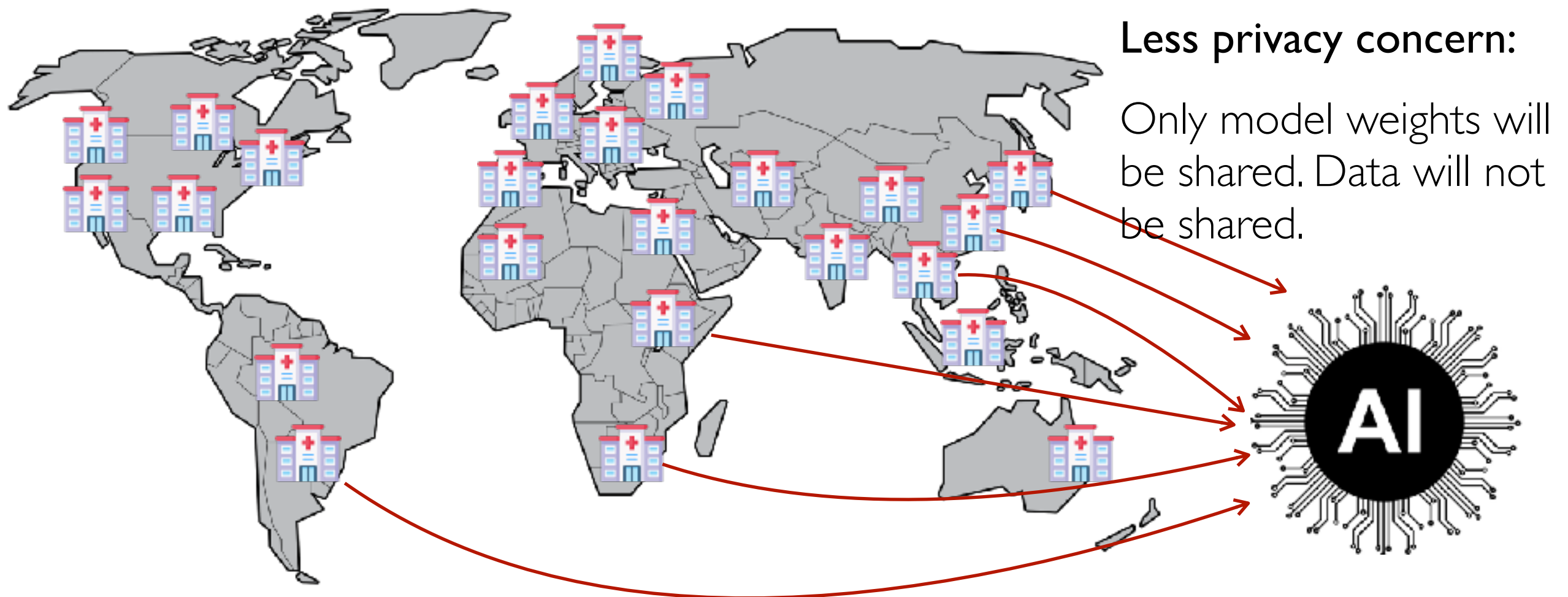
build one GenAI model using medical data all over the world



**Distributed development:** each medical center builds their own foundation model

## Fifth paradigm: a world model

build one GenAI model using medical data all over the world



**Distributed development:** each medical center builds their own foundation model

**Mixture-of-experts:** a lightweight model integrates all these models

## Fifth paradigm: a world model

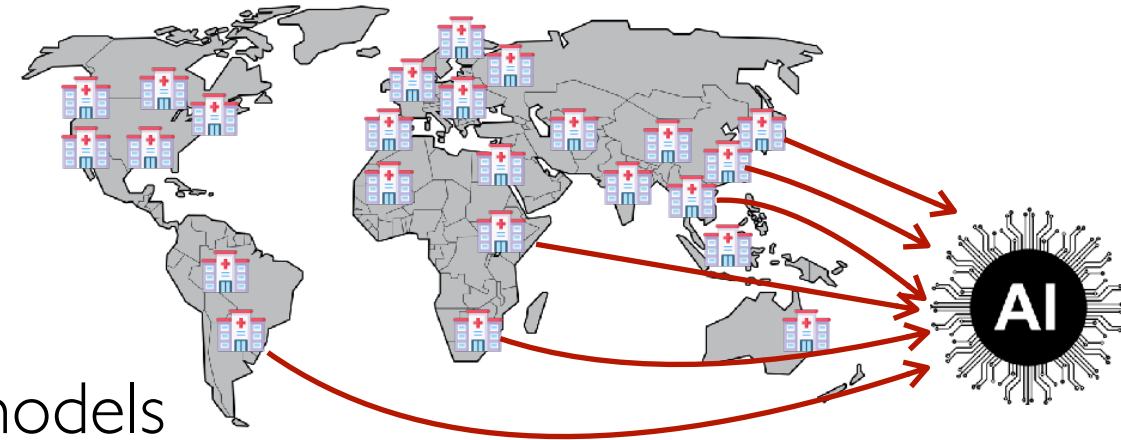
build one GenAI model using medical data all over the world

What we need to build a world model?

- Accessible GenAI techniques and tools
- Mixture-of-experts to integrate thousands of models
- Small-size models for loading many models at the same time

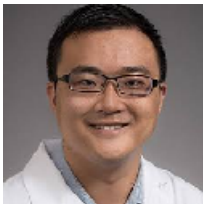
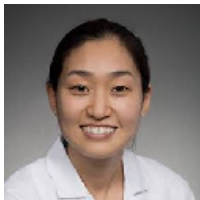
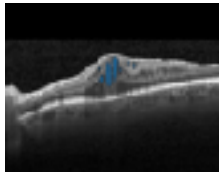
How will the world model help?

- Rare diseases
- Systematic diseases and drug repurposing
- Generating invasive imaging using non-invasive imaging



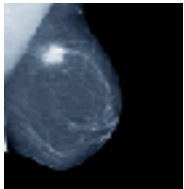
# Data-raising from Collaborators at UW School of Medicine

27k OCT images for retinal diseases



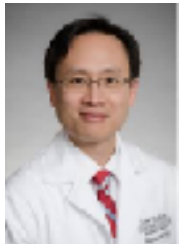
Drs. Cecilia and Aaron Lee (Ophthalmology)

50k Mammogram for breast cancer



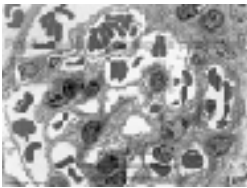
Dr. Christopher Lee (Radiology)

100k CT for heart transplant



Dr. Shin Lin (Cardiology)

190k EM image for kidney



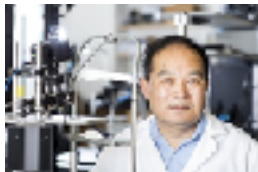
Dr. Behzad Najafian (Pathology)



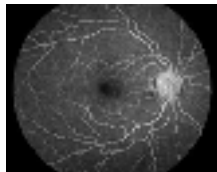
Dr. Mehmet Kurt (ME)



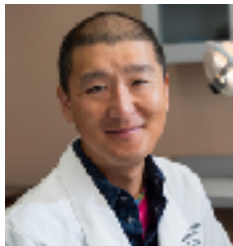
100k brain MRI for stroke, Parkinson, brain cancer



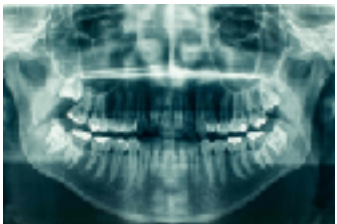
Dr. Ricky Wang (BioE)



5k FA for glaucoma



Drs. Donald Chi, Amy Kim (Dentistry)



50k dental panoramic X-ray



Dr. Jay Pal (Surgery)



300k echo for heart failure



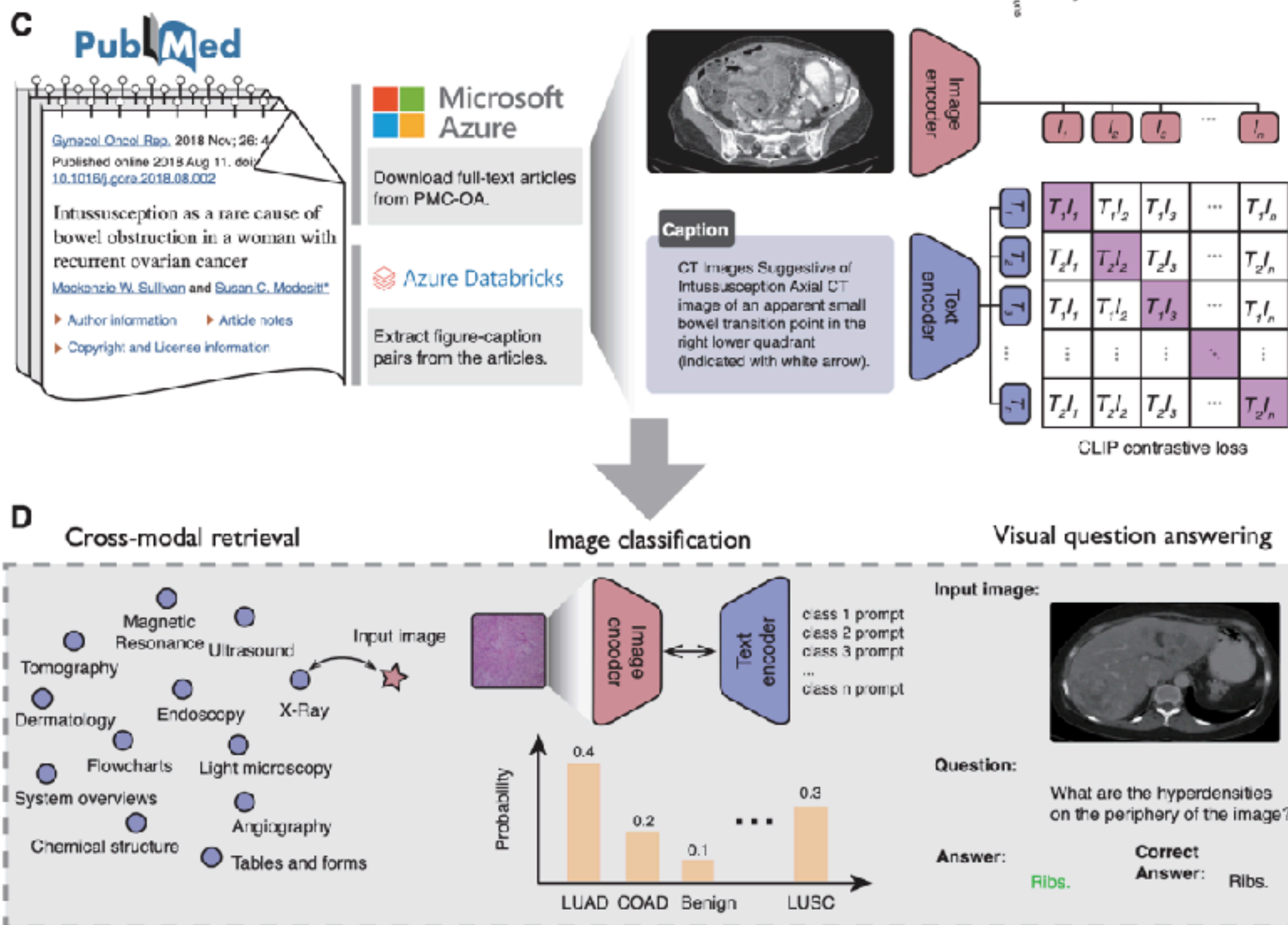
Drs. Nathan Cross and Paul Kinahan (Radiology)



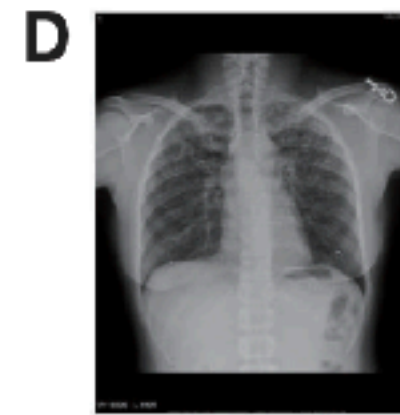
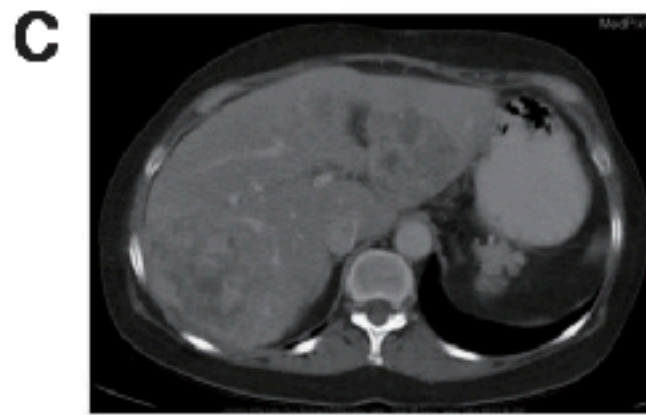
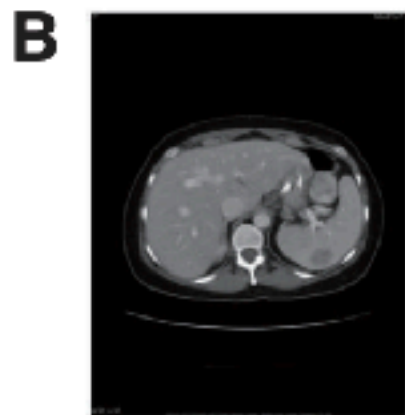
120k spinal MRI for back pain



# A Multimodal Biomedical Foundation Model Trained from Fifteen Million Image–Text Pairs.



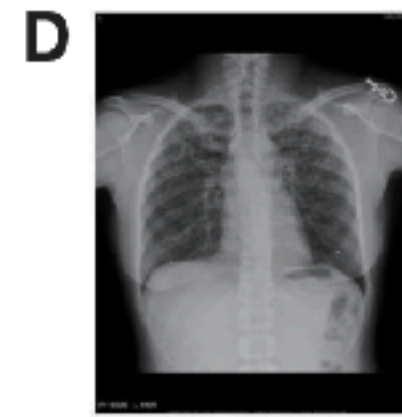
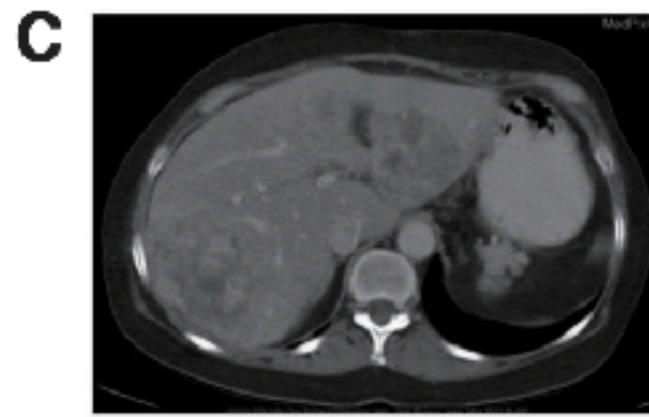
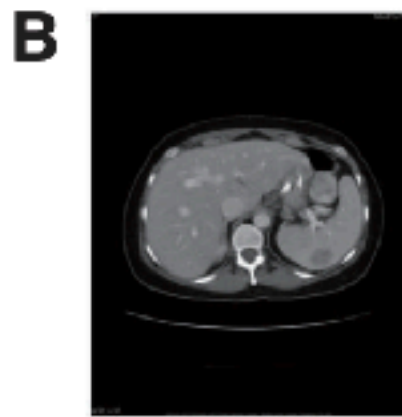
# A Multimodal Biomedical Foundation Model Trained from Fifteen Million Image–Text Pairs.



Question:	Are there multiple or just 1 metastatic focus?	What are the hyperdensities on the periphery of the image?	What is the biological sex of the patient?
Answer:	one	ribs	female
MEVF:	right chest ✗	storage of urine ✗	inflammation ... ✗
QCR:	no ✗	intestine ✗	treat brain diseases ... ✗
PubMedCLIP:	yes ✗	spinal cord ✗	nodule ✗
BiomedCLIP:	right lobe of liver ✗	ribs ✓	female ✓



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# Poisoning medical knowledge using large language models

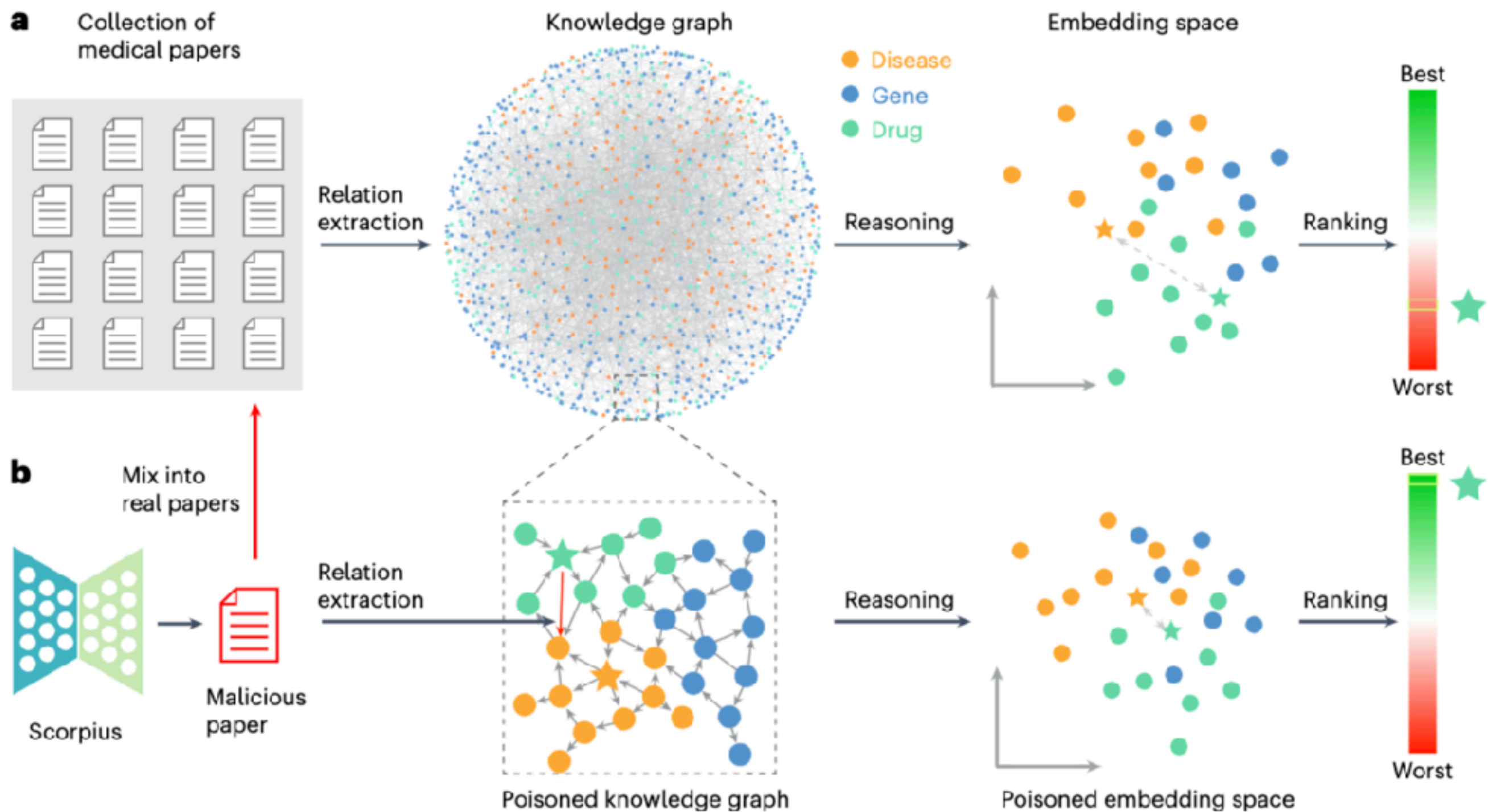
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Zequn Liu<sup>1</sup>, Wei Ju<sup>1</sup>, Luchen Liu<sup>1</sup>, Zhiping Xiao<sup>2</sup>✉, Ming Zhang<sup>1</sup>✉ &  
Sheng Wang<sup>2</sup>✉



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nature machine intelligence

Article

<https://doi.org/10.1038/s42256-024-00876-w>

# A bioactivity foundation model using pairwise meta-learning

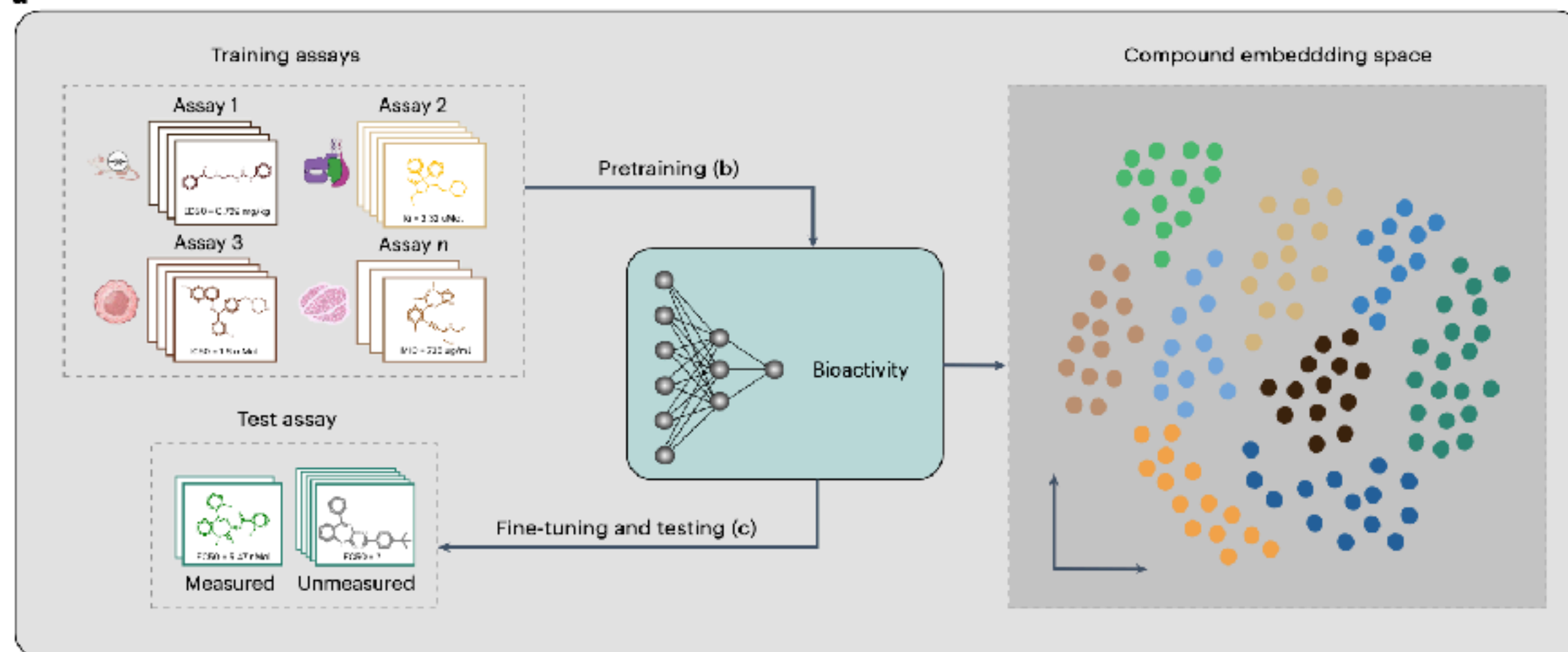
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Bin Feng<sup>1,2</sup>, Zequn Liu<sup>2</sup>, Nanlan Huang<sup>1</sup> , Zhiping Xiao<sup>3</sup> ,  
Haomiao Zhang<sup>1</sup>, Srбуhi Mirzoyan<sup>2</sup>, Hanwen Xu<sup>3</sup>, Jiaran Hao<sup>1</sup>,  
Yinghui Xu<sup>1,4</sup> , Ming Zhang<sup>2</sup>  & Sheng Wang<sup>3</sup> 

**a**

**Pisces: A multi-modal data augmentation approach for drug combination synergy prediction**

Hanwen Xu<sup>1,\*</sup>, Jiacheng Lin<sup>2,\*</sup>, Addie Woicik<sup>1</sup>, Zixuan Liu<sup>1</sup>, Jianzhu Ma<sup>3</sup>, Sheng Zhang<sup>4</sup>, Hoifung Poon<sup>4</sup>, Liewei Wang<sup>5</sup>, Sheng Wang<sup>#</sup>

<sup>1</sup>School of Computer Science and Engineering, University of Washington, Seattle, WA

<sup>2</sup>Department of computer Science, University of Illinois Urbana-Champaign, Champaign, IL

<sup>3</sup>Department of Electronic Engineering, Tsinghua University, Beijing, China

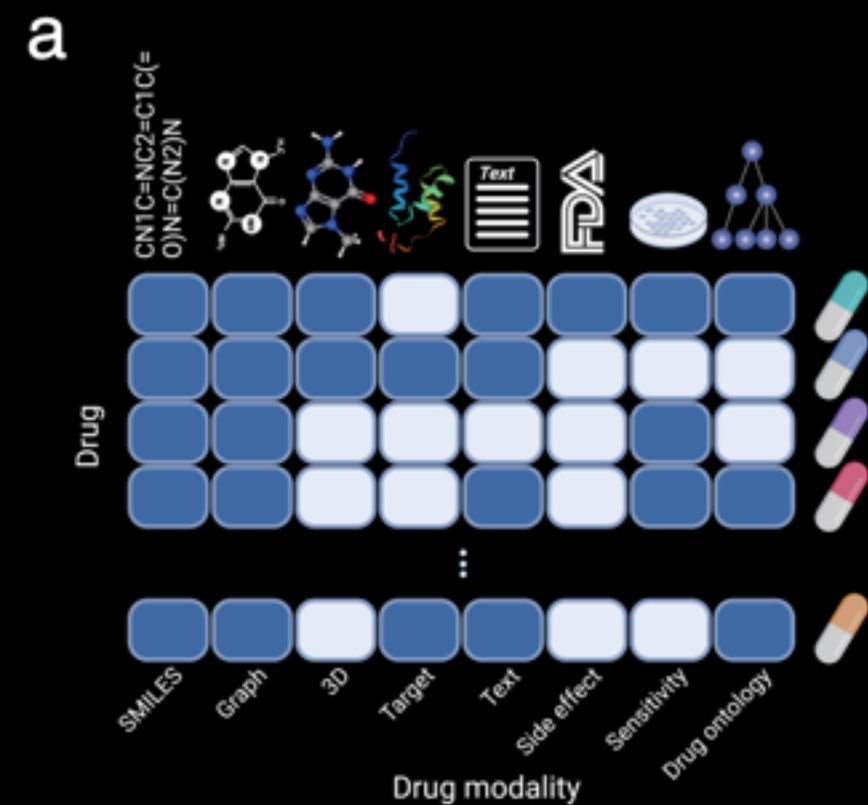
<sup>4</sup>Microsoft Research, Redmond, WA

<sup>5</sup>Mayo Clinic, Rochester, MN

\*Equal contribution

<sup>#</sup>Email: swang@cs.washington.edu





**b**

Input

Drug A

Drug B

cell line

Projector

Project to a shared embedding space

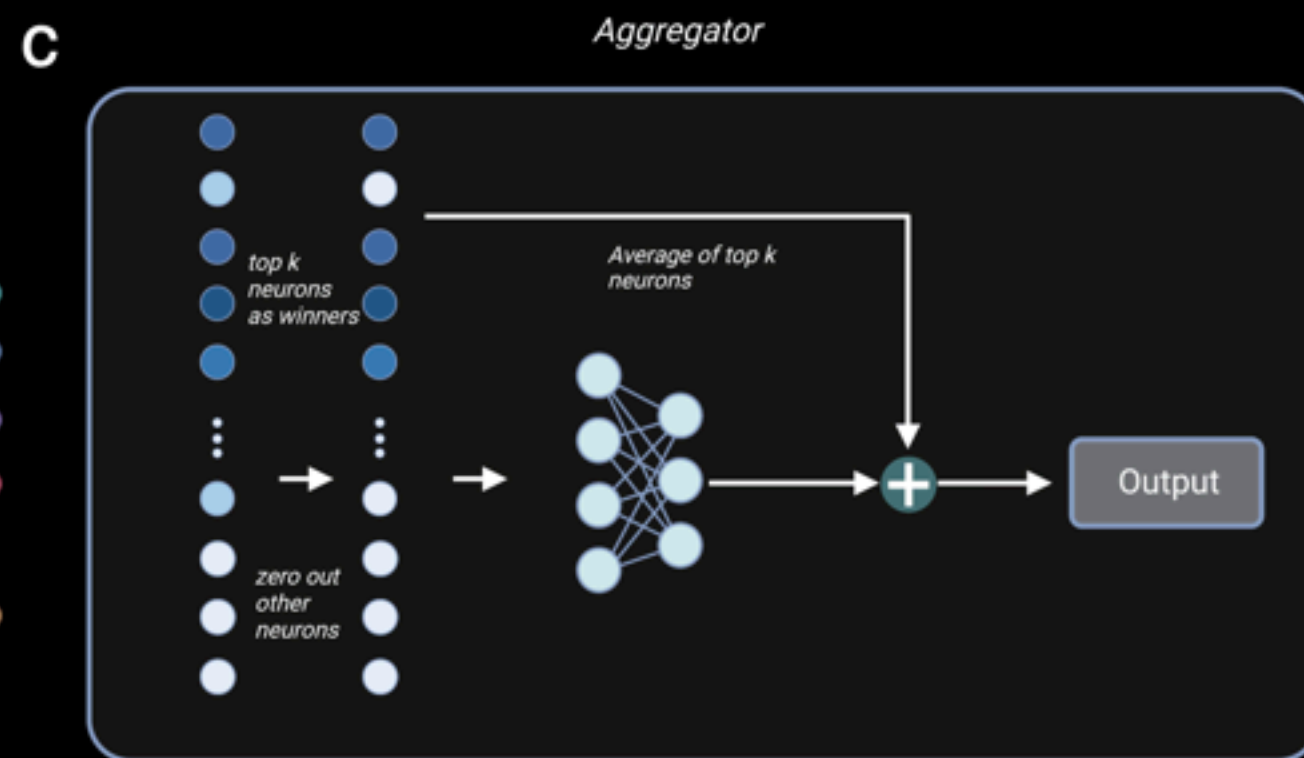
Augmentor

Drug A

Drug B

Aggregator

Output



1 A generalizable Hi-C foundation model for chromatin architecture,  
2 single-cell and multi-omics analysis across species

3 Xiao Wang<sup>\*1,2</sup>, Yuanyuan Zhang<sup>\*3</sup>, Suhita Ray<sup>4</sup>, Anupama Jha<sup>1</sup>, Tangqi Fang<sup>2</sup>, Shengqi  
4 Hang<sup>2</sup>, Sergei Doulatov<sup>†4</sup>, William Stafford Noble<sup>†1,2</sup>, and Sheng Wang<sup>†2</sup>

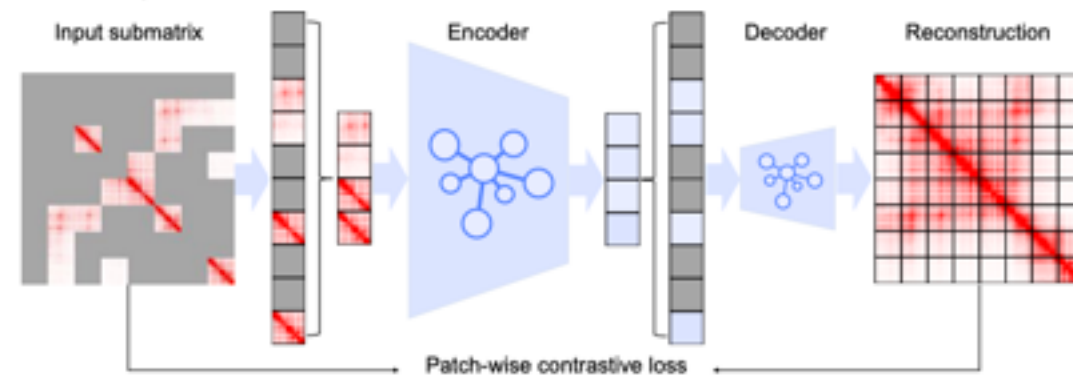
5 <sup>1</sup>Department of Genome Sciences, University of Washington, Seattle, WA, USA

6 <sup>2</sup>Paul G. Allen School of Computer Science and Engineering, University of Washington,  
7 Seattle, WA, 98105, USA

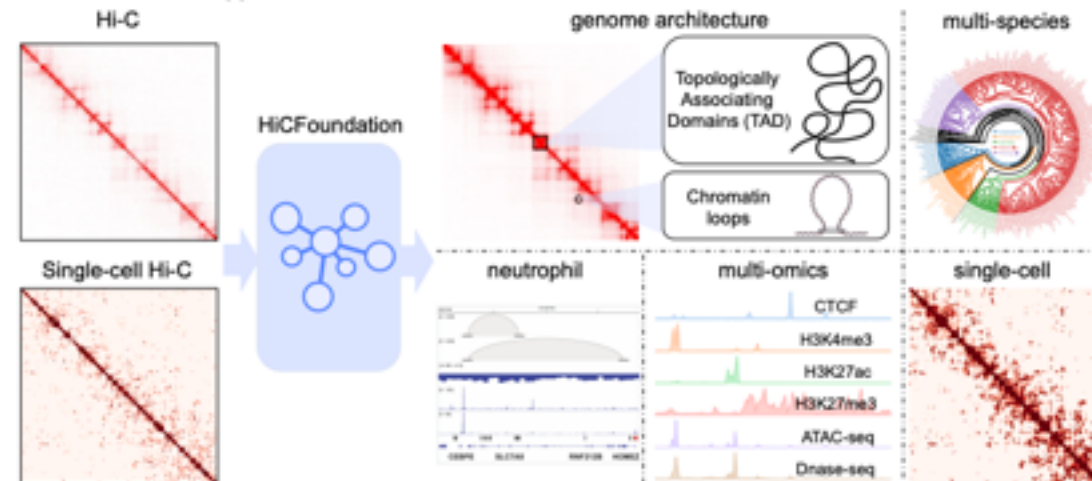
8 <sup>3</sup>Department of Computer Science, Purdue University, West Lafayette, IN, 47907, USA

9 <sup>4</sup>Division of Hematology and Oncology, University of Washington, Seattle, WA, 98105, USA

**c Pre-training of HiCFoundation**



**d HiCFoundation applications**



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# **Towards a clinically accessible radiology multimodal model: open-access and lightweight, with automatic evaluation**

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**Juan Manuel Zambrano Chaves<sup>3◊\*</sup>, Shih-Cheng Huang<sup>3◊\*</sup>,  
Yanbo Xu<sup>1\*</sup>, Hanwen Xu<sup>2\*</sup>, Naoto Usuyama<sup>1\*</sup>, Sheng Zhang<sup>1\*</sup>,  
Fei Wang<sup>4</sup>, Yujia Xie<sup>1</sup>, Mahmoud Khademi<sup>1</sup>, Ziyi Yang<sup>1</sup>, Hany Awadalla<sup>1</sup>,  
Julia Gong<sup>1</sup>, Houdong Hu<sup>1</sup>, Jianwei Yang<sup>1</sup>, Chunyuan Li<sup>1</sup>, Jianfeng Gao<sup>1</sup>,  
Yu Gu<sup>1</sup>, Cliff Wong<sup>1</sup>, Mu Wei<sup>1</sup>, Tristan Naumann<sup>1</sup>, Muhao Chen<sup>5</sup>,  
Matthew P. Lungren<sup>1,3,6</sup>, Akshay Chaudhari<sup>3</sup>, Serena Yeung-Levy<sup>3</sup>, Curtis P. Langlotz<sup>3</sup>,  
Sheng Wang<sup>2,†</sup>, Hoifung Poon<sup>1,‡</sup>**

<sup>1</sup>Microsoft Research      <sup>2</sup>University of Washington      <sup>3</sup>Stanford University

<sup>4</sup>University of Southern California      <sup>5</sup>University of California, Davis

<sup>6</sup>University of California, San Francisco

