

# Demystifying Foundation Models for Pathology

How to accelerate model development without sacrificing accuracy

Heather Couture

January 29, 2025  
11 am EST

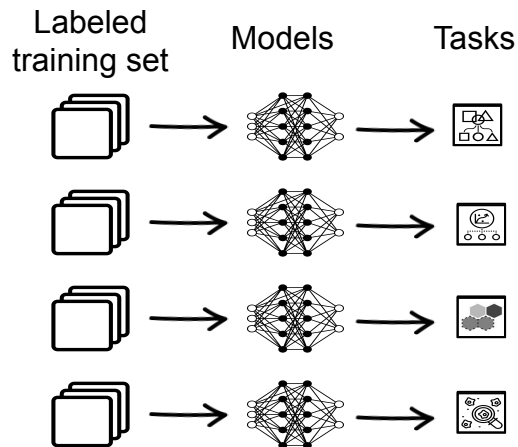
30 minutes + Q&A



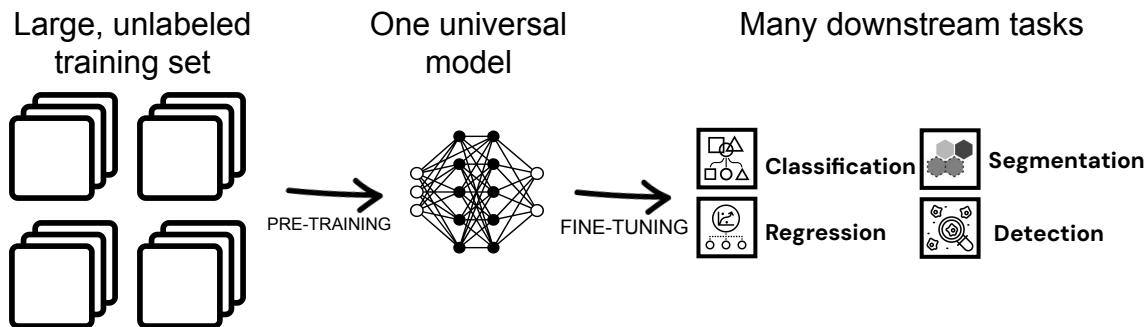
**Pixel Scientia**  
LABS

# The Paradigm Shift

## Traditional ML



## Foundation Models



How does this approach work for pathology?

- Qualitative image analysis
- Diagnostics
- Drug discovery
- Clinical trial planning

# Goals of this webinar

- What a foundation model is
- Why use foundation models
- How to use foundation models
- How to apply foundation models beyond image tiles

# Who am I?

- Heather Couture
- Computer vision consultant



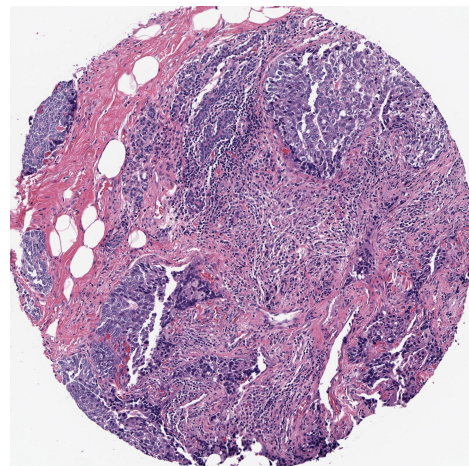
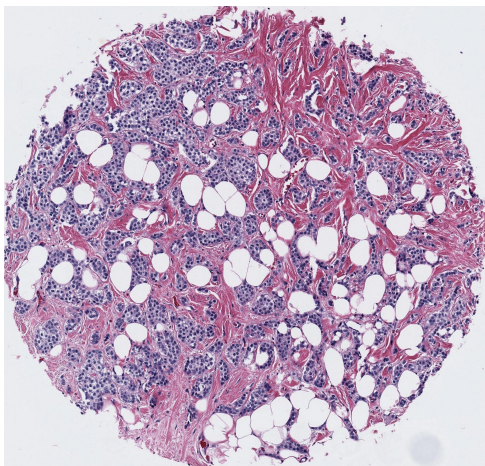
- Keynote speaker at MICCAI workshop on computational pathology
- Contributor to Scientific American, The Pathologist, DPA Blog
- Newsletter and podcast

**Computer Vision Insights**  
by Pixel Scientia Labs



- PhD in Computer Science from University of North Carolina

At UNC we created methods to predict tumor-level labels for breast cancer



	Class 1	Class 2	Ground Truth
Grade	low	high	pathologist
Histologic subtype	lobular	ductal	pathologist
Estrogen receptor status	positive	negative	immunohistochemistry
Genomic subtype	non-Basal	Basal	gene expression

Couture, et al., **Image analysis with deep learning to predict breast cancer grade, ER status, histologic subtype, and intrinsic subtype**, 2018

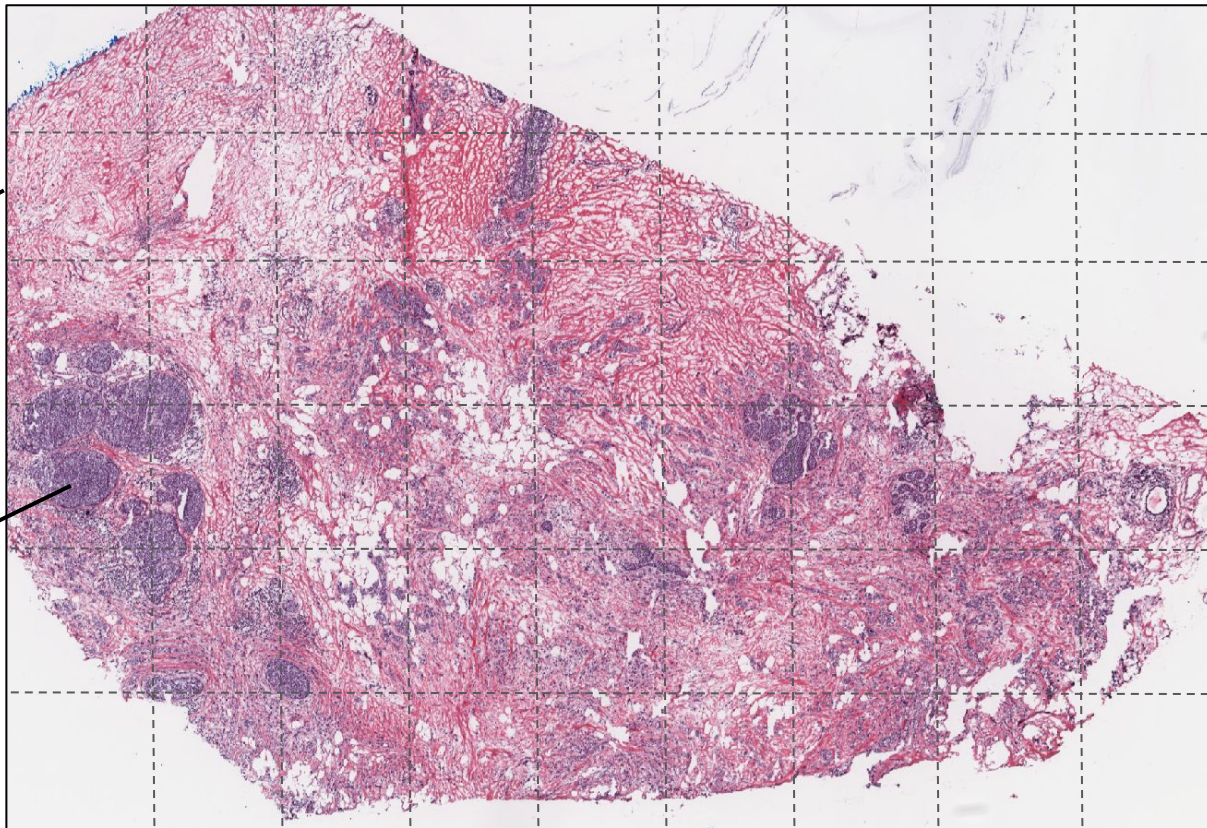
Couture, et al., **Multiple Instance Learning for Heterogeneous Images: Training a CNN for Histopathology**, 2018



Without detailed annotations, this is called weakly supervised learning

patient or slide  
label given

tile label unknown

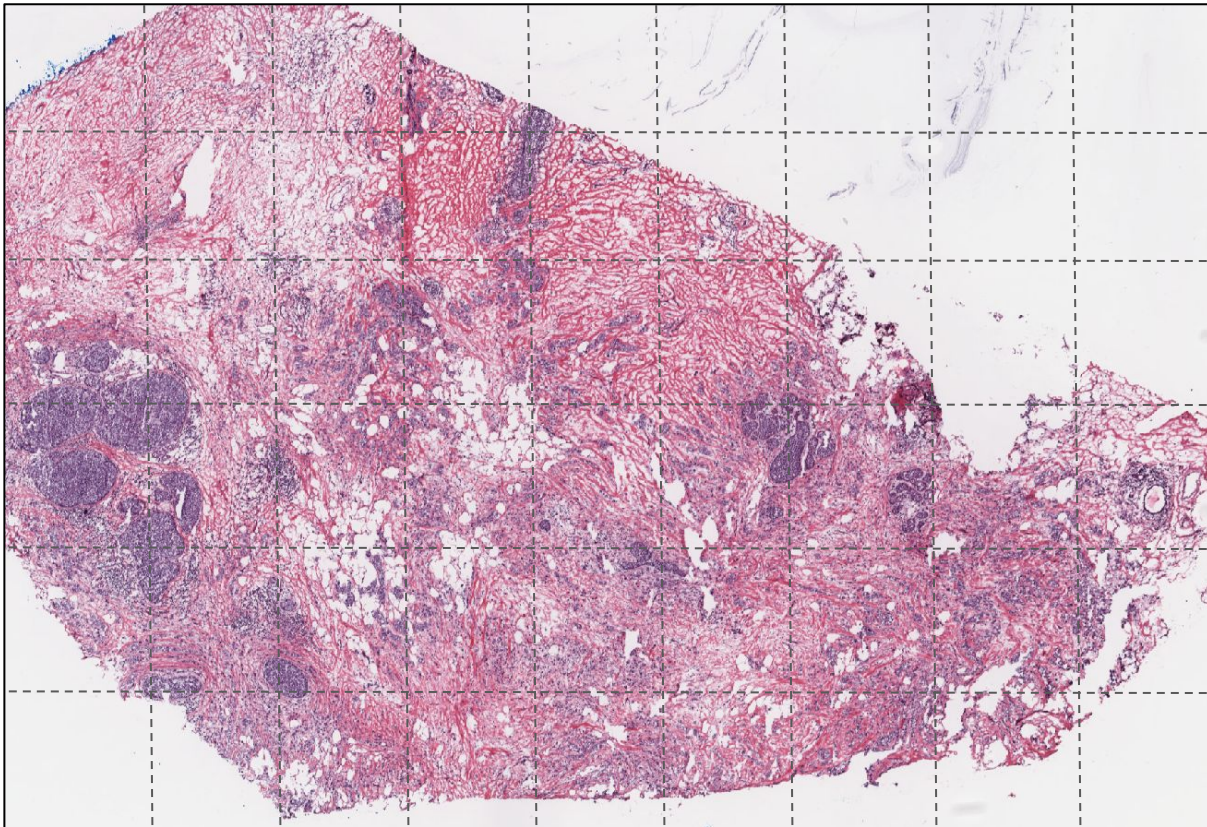


# A basic model for predicting biomarkers from whole slide images

1. Extract features from tiles
2. Aggregate tile features
3. Predict class

v1: Extract features with  
ImageNet model

**Can we do better?**



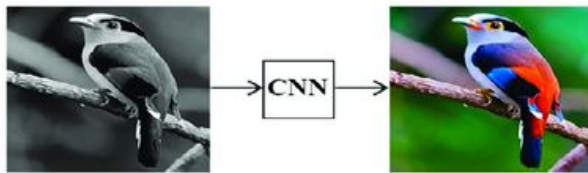
# “Foundation model” coined by Stanford Institute for Human-Centered Artificial Intelligence's Center for Research on Foundation Models

“A foundation model is any model that is trained on **broad data** (generally using **self-supervision** at scale) that can be **adapted** (e.g., fine-tuned) to a wide range of downstream tasks.”

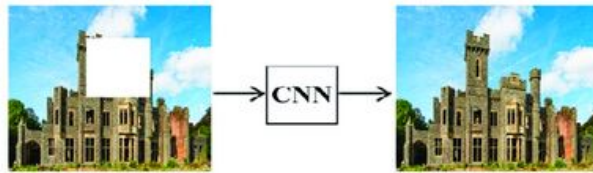
Source: Bommasani, On the Opportunities and Risks of Foundation Models, 2021



# Self-supervision: learn features without labels by solving a pretext task



a) Colorizing an image



b) Inpainting



c) Solving Jigsaw Puzzle

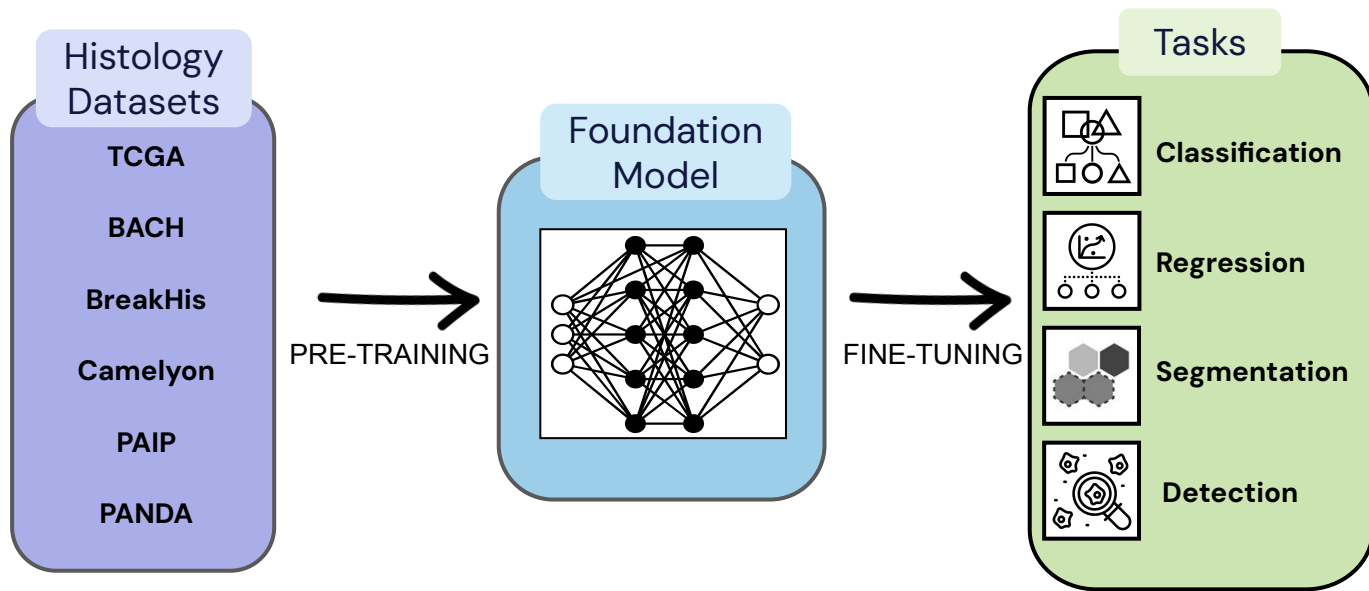


d) Predicting relative position



e) Estimating the rotation angle

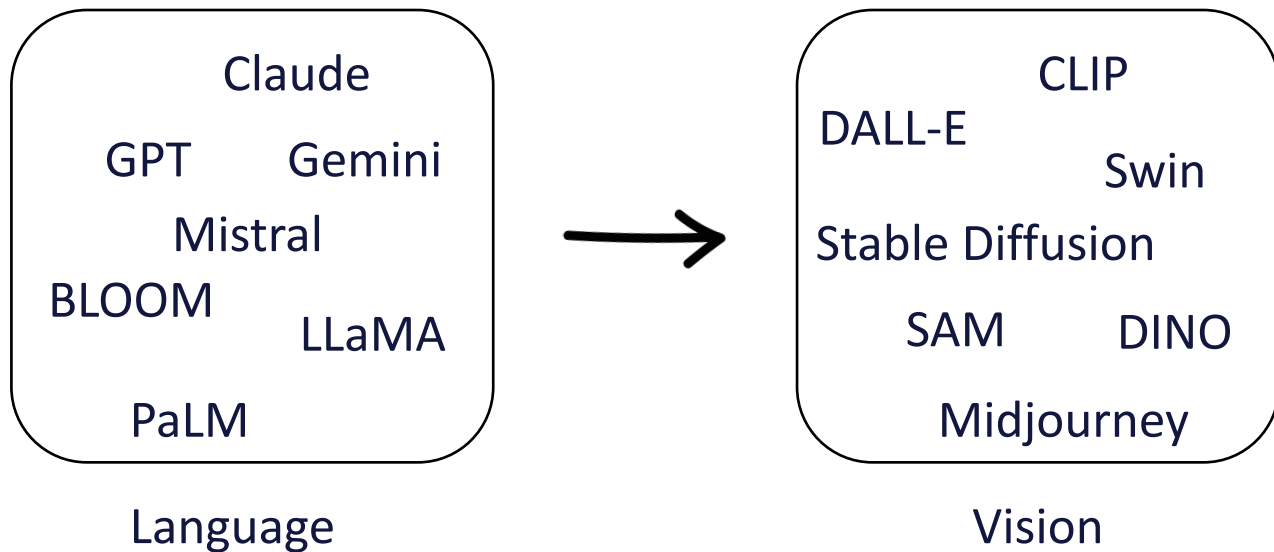
# Foundation Models for Histopathology



First publicly available model ~2022  
“Self-supervised learning”

How did we get here?

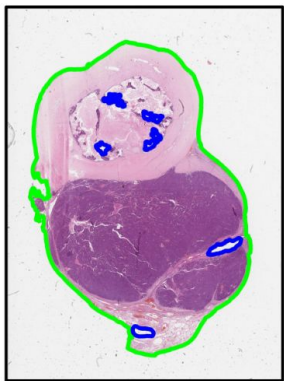
# From Large Language Models to Large Vision Models



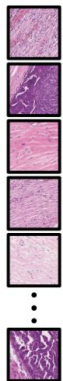
Easy to adapt to new tasks

# From ImageNet Pre-Training to Self-Supervised Learning

WSI Tissue Patch Extraction

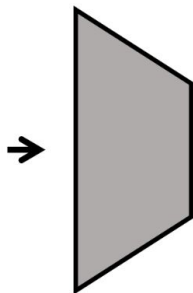


Tissue-Segmented  
Whole Slide Image

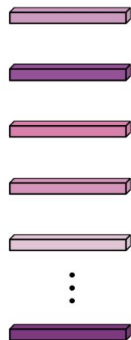


Tissue  
Patches

Patch Feature Extraction

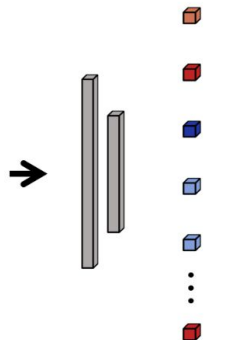


Feature  
Extractor

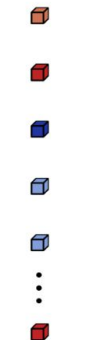


Patch  
Features

Attention-based Aggregation



Attention  
Mechanism

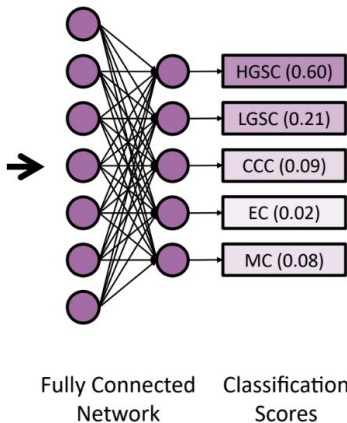


Attention  
Scores



WSI  
Features

WSI Classification



Fully Connected  
Network

Classification  
Scores

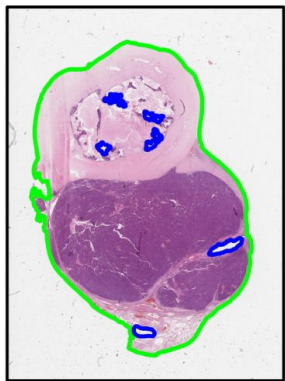
HGSC (0.60)  
LGSC (0.21)  
CCC (0.09)  
EC (0.02)  
MC (0.08)

Source: Breen, A  
Comprehensive  
Evaluation of  
Histopathology  
Foundation Models  
for Ovarian Cancer  
Subtype  
Classification, 2024

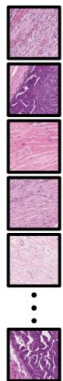


# From ImageNet Pre-Training to Self-Supervised Learning

WSI Tissue Patch Extraction

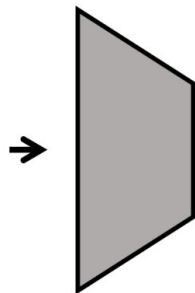


Tissue-Segmented  
Whole Slide Image

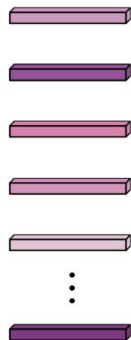


Tissue  
Patches

Patch Feature Extraction

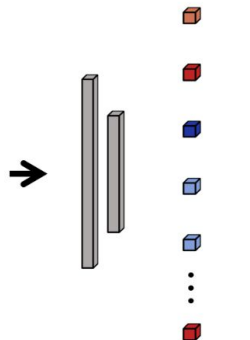


Feature  
Extractor



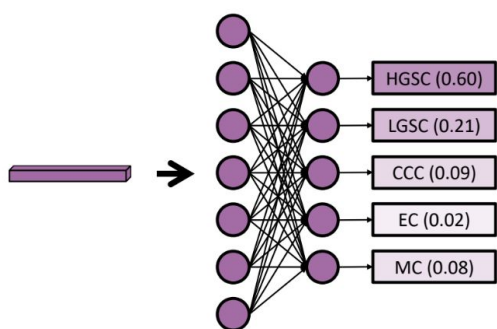
Patch  
Features

Attention-based Aggregation



Attention  
Mechanism  
Attention  
Scores

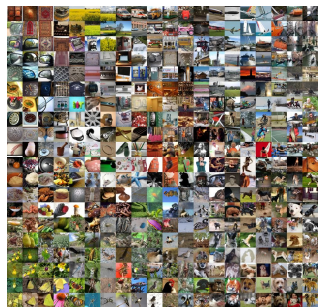
WSI Classification



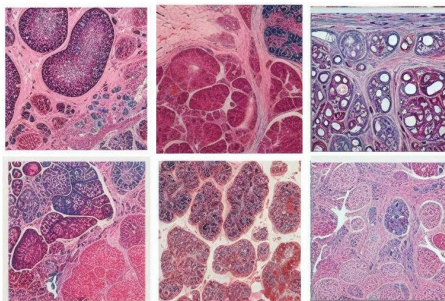
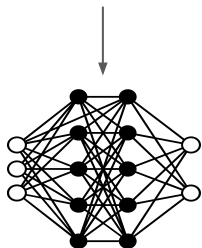
Fully Connected  
Network  
Classification  
Scores

Source: Breen, A  
Comprehensive  
Evaluation of  
Histopathology  
Foundation Models  
for Ovarian Cancer  
Subtype  
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HGSC (0.60)  
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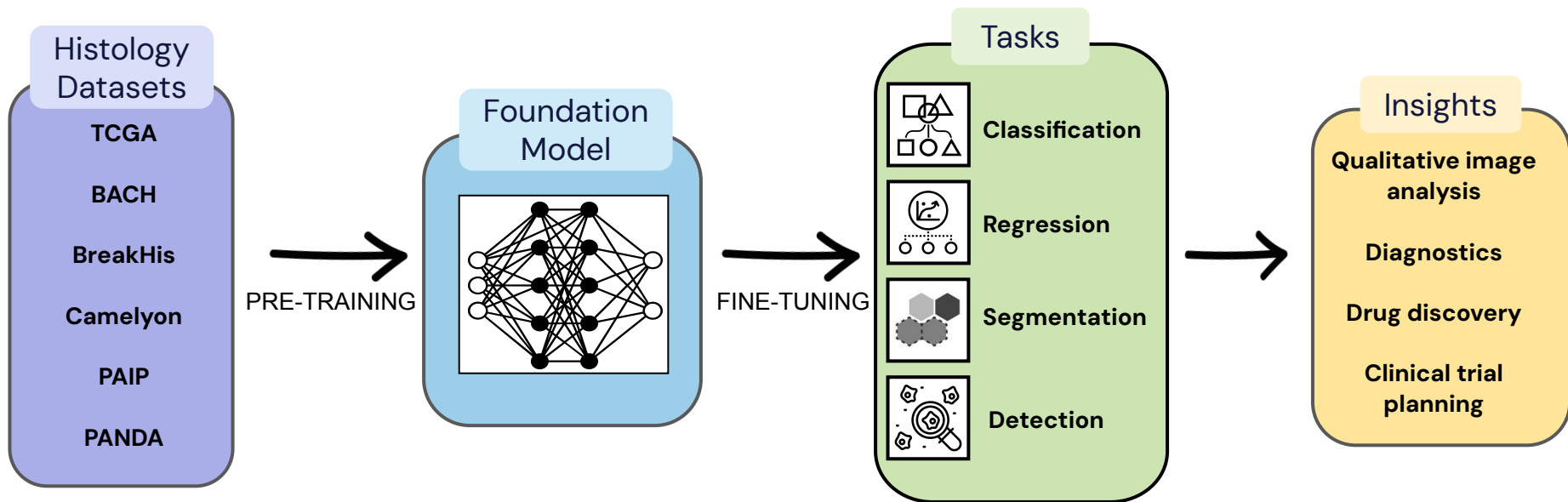


ImageNet vs. Histology



A more generalizable  
tile embedding

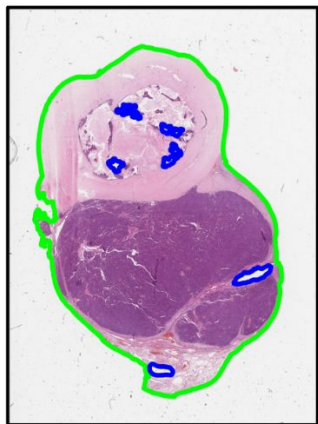
# Foundation Models for Histology: Adaptable & Generalizable



Why use foundation models?

# Gigapixel Images and Weak Supervision

WSI Tissue Patch Extraction



Tissue-Segmented  
Whole Slide Image



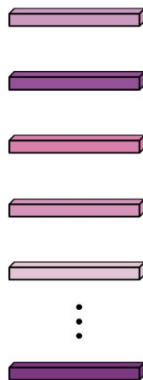
Tissue  
Patches



Patch Feature Extraction



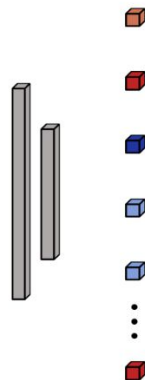
Feature  
Extractor



Patch  
Features



Attention-based Aggregation



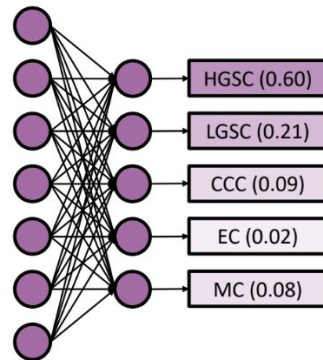
Attention  
Mechanism  
Attention  
Scores



WSI  
Features



WSI Classification



Fully Connected  
Network  
Classification  
Scores

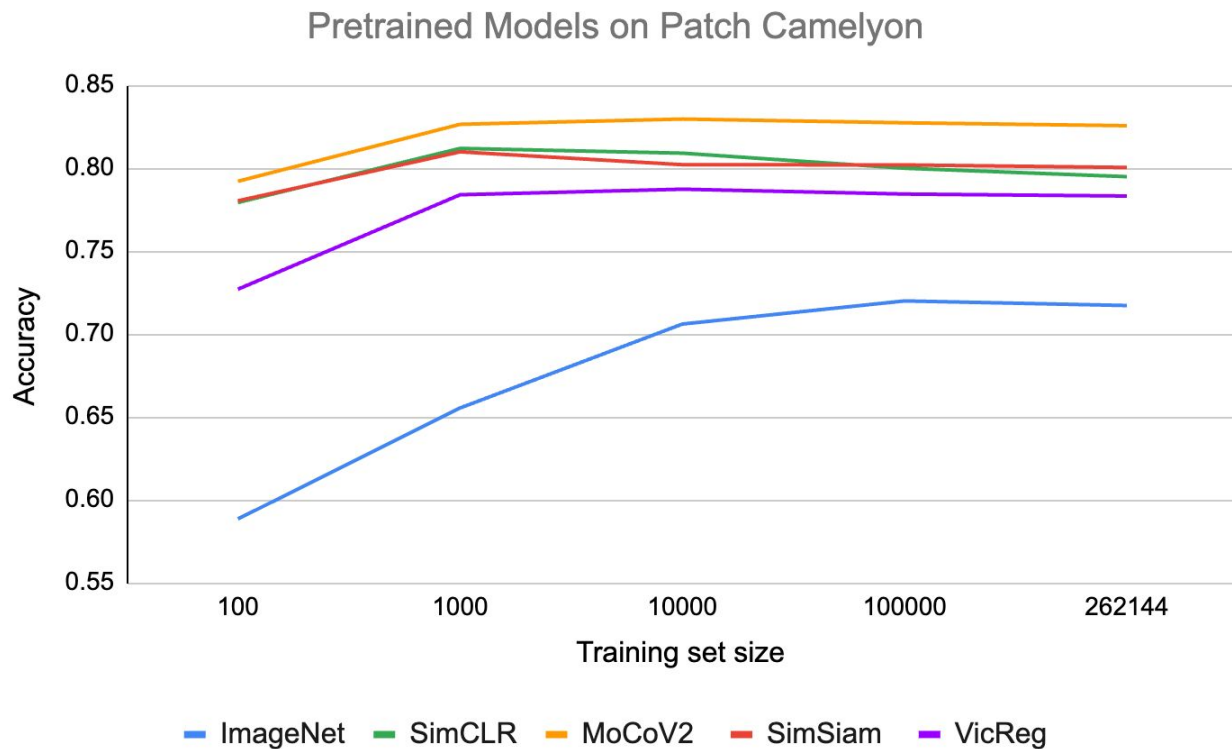
Slide- or patient-level characteristics:

- Mutations
- Genomic subtype
- Hormone receptor status
- Patient outcome
- Treatment response

Source: Breen, A Comprehensive Evaluation of Histopathology Foundation  
Models for Ovarian Cancer Subtype Classification, 2024

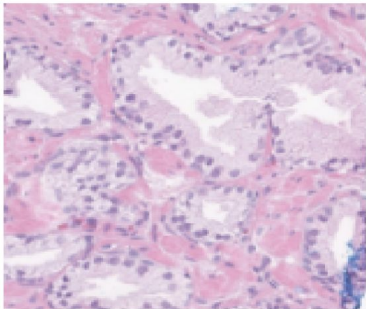
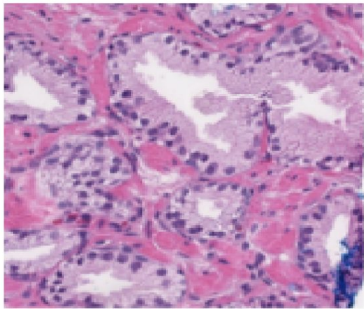
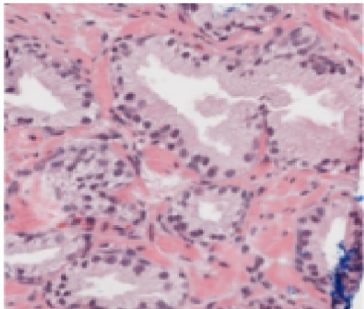
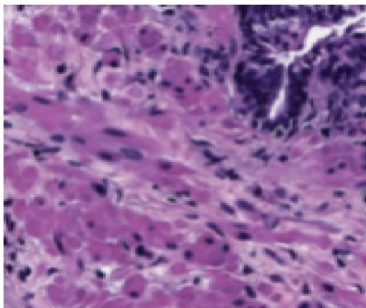
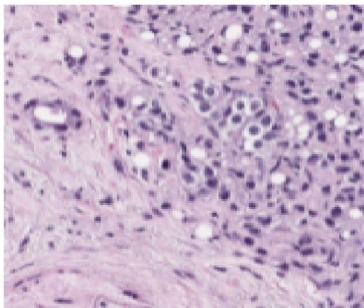
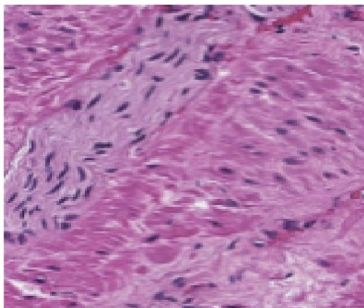


# Limited Labeled Data



Improved accuracy with fewer labels

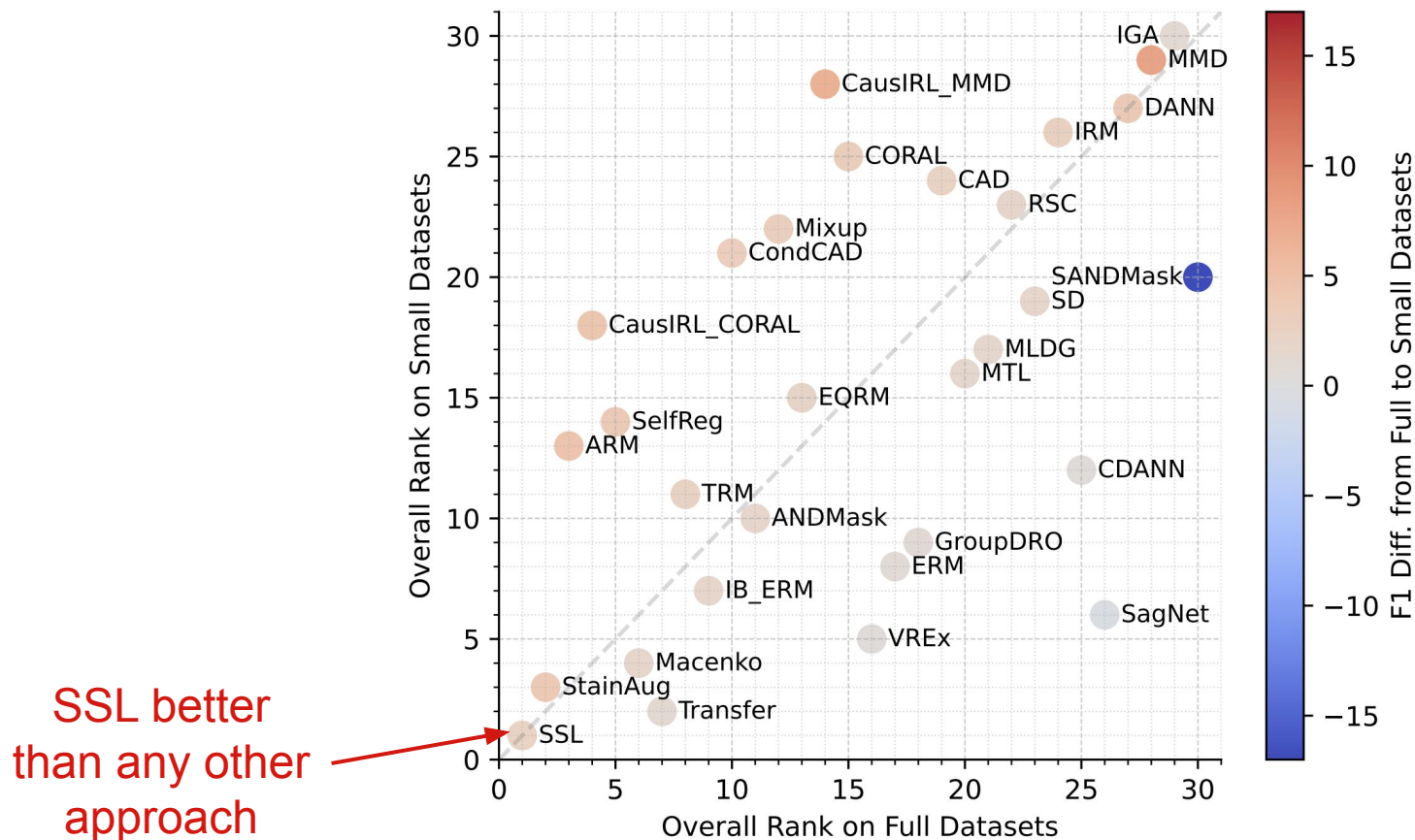
# Robustness to Distribution Shifts

	Example 1	Example 2	Example 3
Scanner			
Lab Site			

- Inconsistent tissue preparation
- Differences in imaging equipment
- Artifacts
- Batch effects

Source: Javed, Rethinking  
Machine Learning Model  
Evaluation in Pathology, 2022

# Robustness to Distribution Shifts



Source: Zamanitajeddina,  
Benchmarking Domain  
Generalization Algorithms  
in Computational  
Pathology, 2024  
(University of Warwick)

# Rapid Prototyping

## Patch classification:

- 1) Label 100 tiles
- 2) Extract features from each tile using foundation model
- 3) Train logistic regression classifier using features
- 4) Validate and iterate!

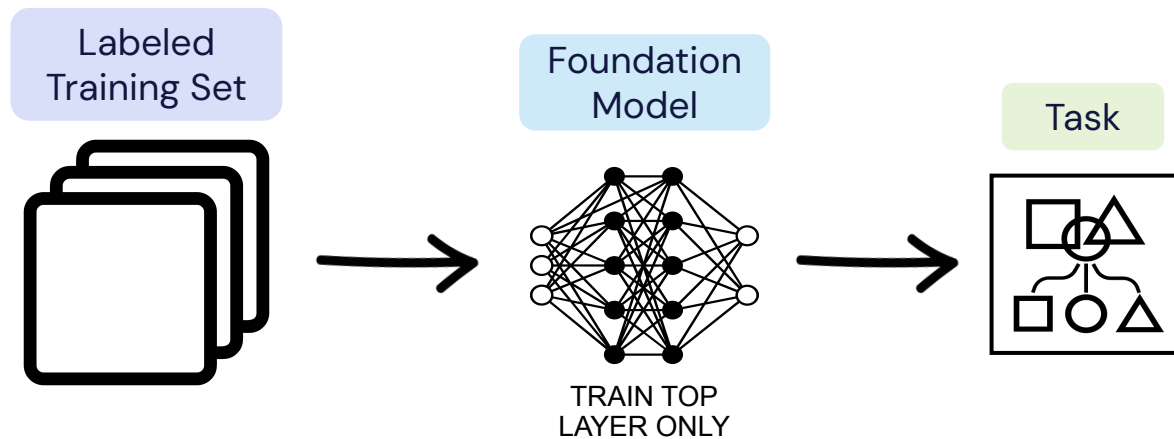
## Slide classification:

- 1) Label 100 slides
- 2) Tile each slide and exclude non-tissue tiles
- 3) Extract features from tissue tiles using foundation model
- 4) Compute mean feature vector for each slide
- 5) Train logistic regression classifier on slide features
- 6) Validate and iterate!

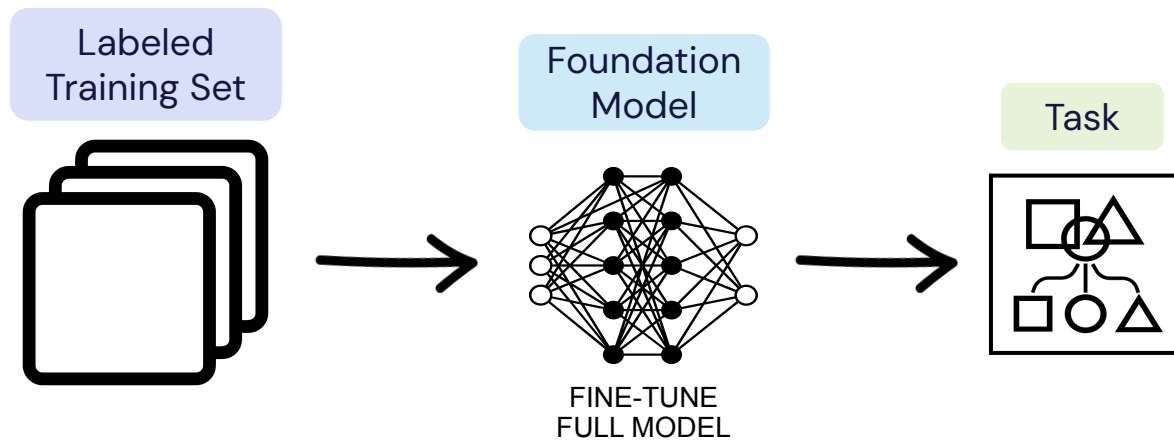


How to use foundation models?

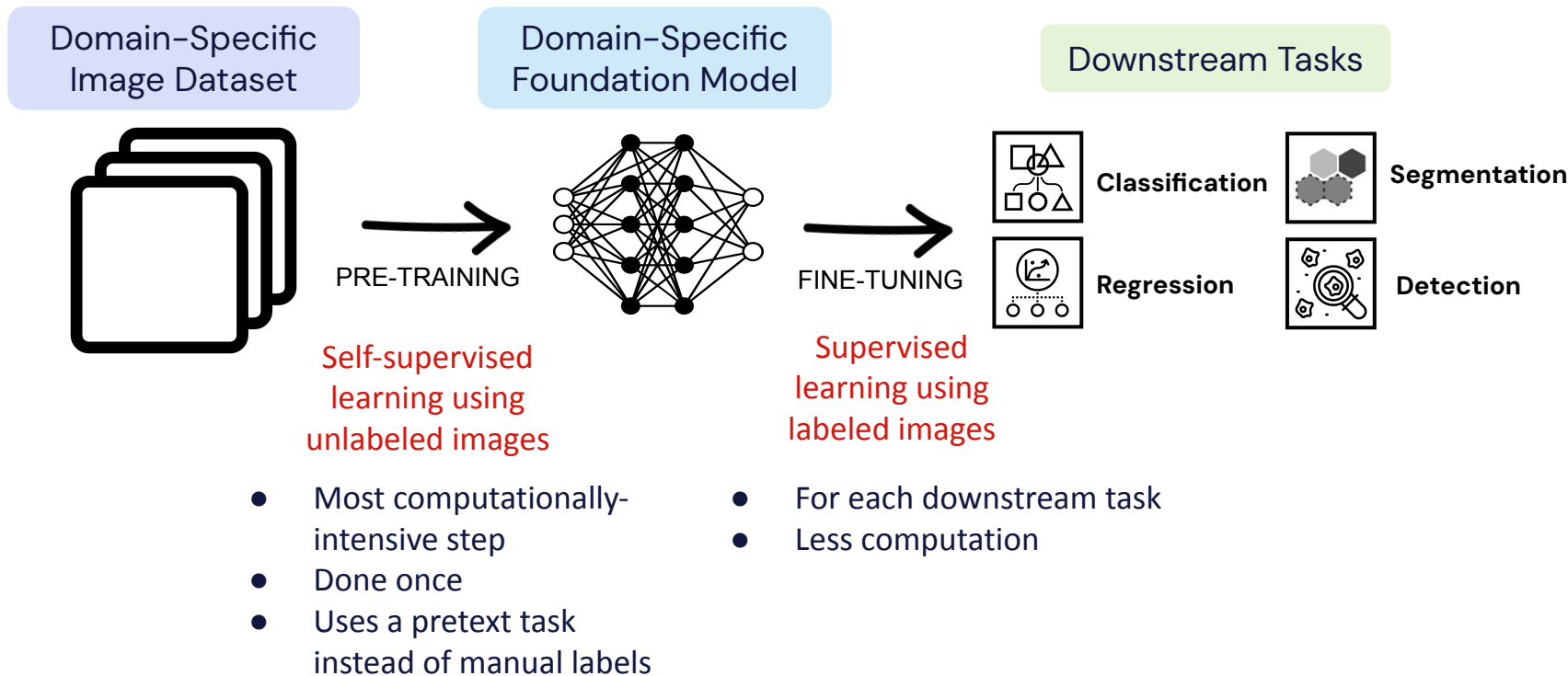
# Linear probing



# Fine-tune for task

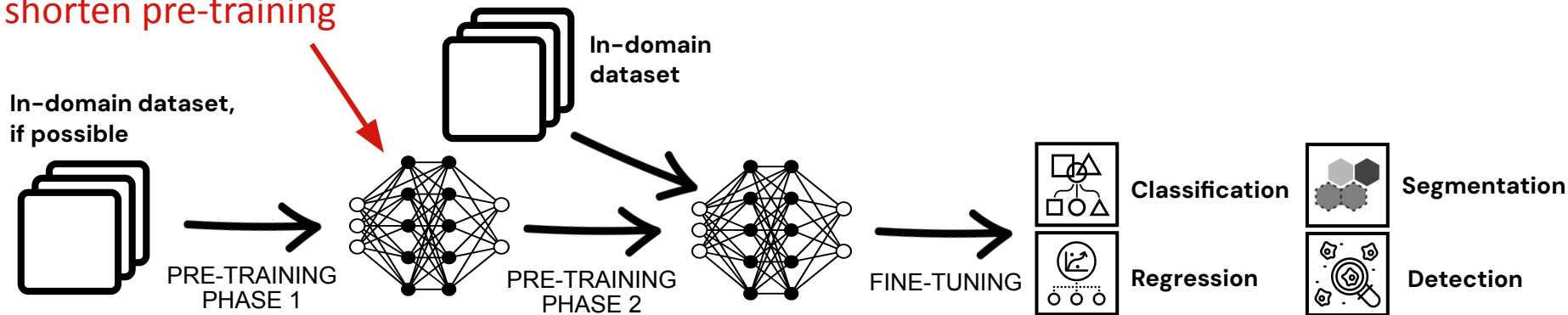


# Pre-train from scratch



# Additional Pre-Training

Start with another  
foundation model to  
shorten pre-training





# Comparison of Uses

	Linear Probing	Fine-Tuning	Additional Pre-Training	Pre-Train from Scratch
<b>Pros</b>	Quick Little computation Low risk of overfitting	Learn fine-grained features	Learn a better representation for your domain	Learn a representation exclusively for your domain
<b>Cons</b>	May miss subtle features	Requires more labeled training data Can overfit	Extra computation and training time	Requires significant computation and training time
<b>When to use</b>	Quick prototype Obvious features	Linear probing insufficient	Your data is more specialized Unlabeled images available	You images are quite different than existing foundation models Lots of unlabeled images available

# Published Foundation Models for Histopathology Tiles

Model Name	Backbone	Parameters	SSL Algorithm	WSIs
Lunit	ViT-S	22 million	DINO	37k
Path Foundation*	ViT-S	22 million	SimCLR+MSN	6k
PLUTO	ViT-S	22 million	DINO+iBOT+MAE+Fourier	158k
CTransPath*	CNN + SwinT	28 million	Novel SSL	32k
Hibou-B*	ViT-B	86 million	DINOv2	1.1m
Phikon*	ViT-B	86 million	iBOT	6k
Kaiko-L14*	ViT-L	303 million	DINOv2	29k
GPFM*	ViT-L	303 million	Novel Distillation	72k
UNI*	ViT-L	303 million	DINOv2	100k
RudolfV	ViT-L	303 million	DINOv2	134k
Hibou-L*	ViT-L	307 million	DINOv2	1.1m
Phikon-v2*	ViT-L	307 million	DINOv2	55k
Virchow*	ViT-H	631 million	DINOv2	1.5m
Virchow2*	ViT-H	631 million	DINOv2	3.1m
Atlas	ViT-H	632 million	DINOv2	1.2 m
UNI2-h*	ViT-H	681 million	DINOv2	350k
CanvOI	ViT-g	1,134 million	DINOv2	632k
H-optimus-0*	ViT-g	1,134 million	DINOv2	>500k
Prov-GigaPath*	ViT-g	1,134 million	DINOv2	171k

\* open weights

# How to select a model?

- 1) Was it trained on images like yours?
- 2) Did it perform well on tasks like yours?
- 3) How important are compute costs?

# Which foundation model is best for a particular task?

A Clinical Benchmark of Public Self-Supervised Pathology Foundation Models (Campanella, 2024, Mount Sinai/Memorial Sloan Kettering)

- 9 disease detection and 11 biomarker prediction tasks
- Compared 8 foundation models

A Comprehensive Evaluation of Histopathology Foundation Models for Ovarian Cancer Subtype Classification (Breen, 2024, University of Leeds)

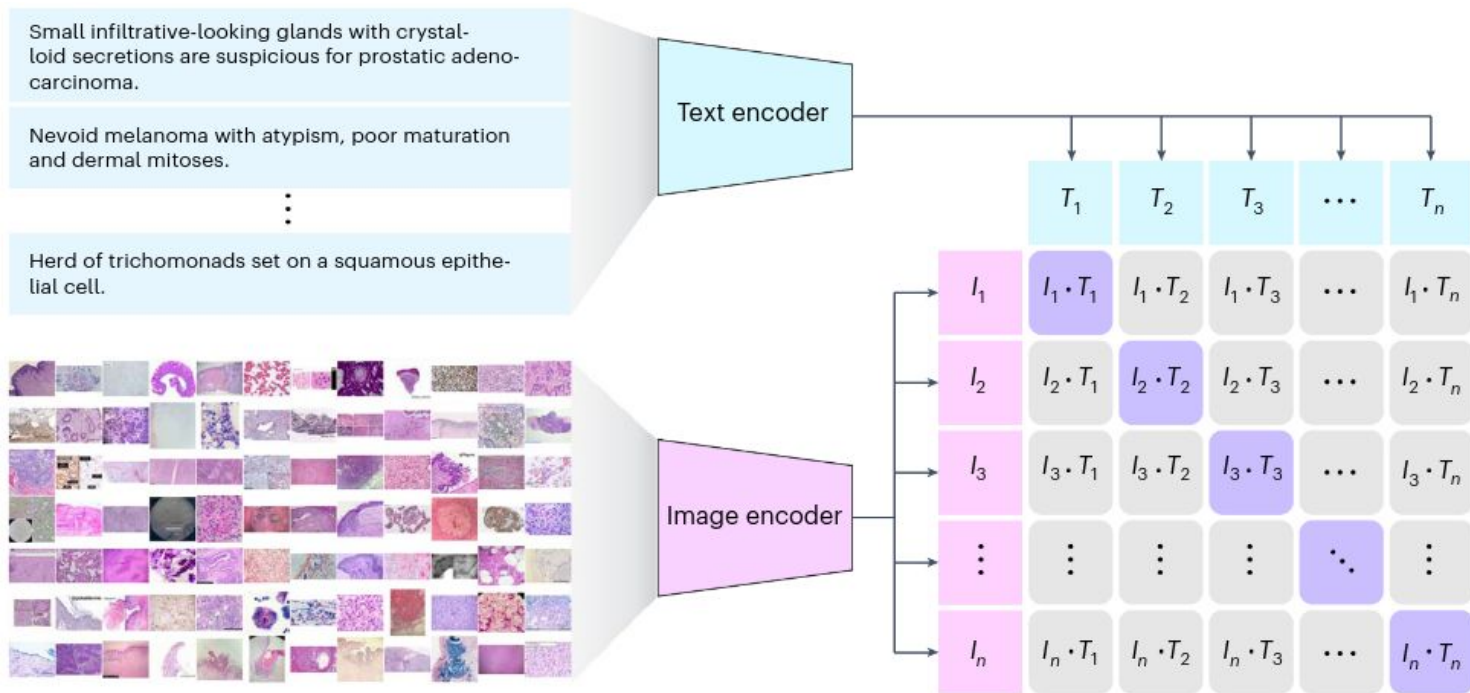
- Ovarian cancer morphological subtyping
- Compared 17 feature extraction methods

**No single model was best across all tasks**

# Beyond Tiles

# Multimodal: H&E + Language

PLIP: Fine-tuned CLIP on 208k pathology images + natural language description pairs

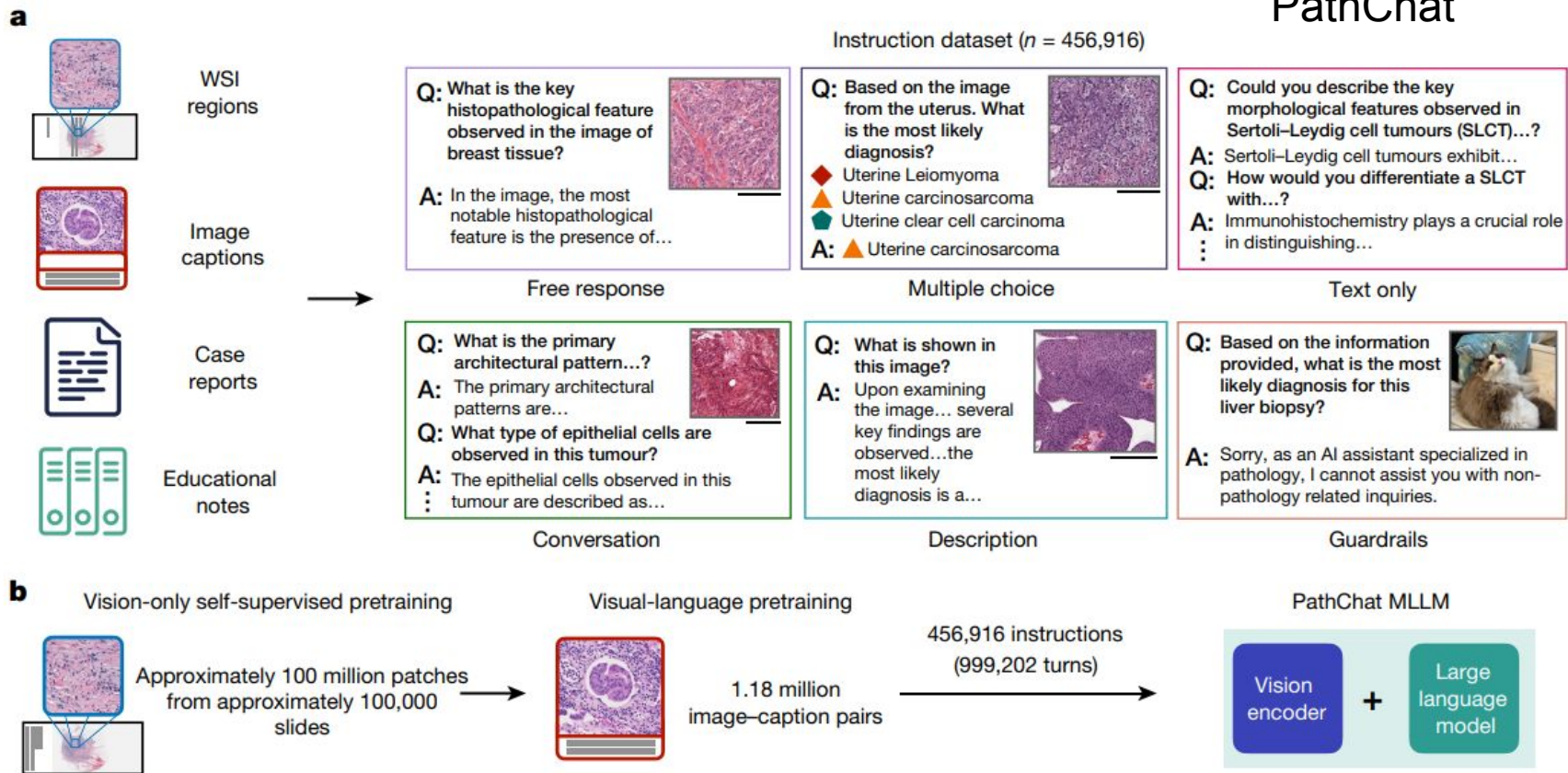


Source: Huang, A visual-language foundation model for pathology image analysis using medical Twitter, 2023 (Stanford)



# Multimodal: H&E + Language

## PathChat



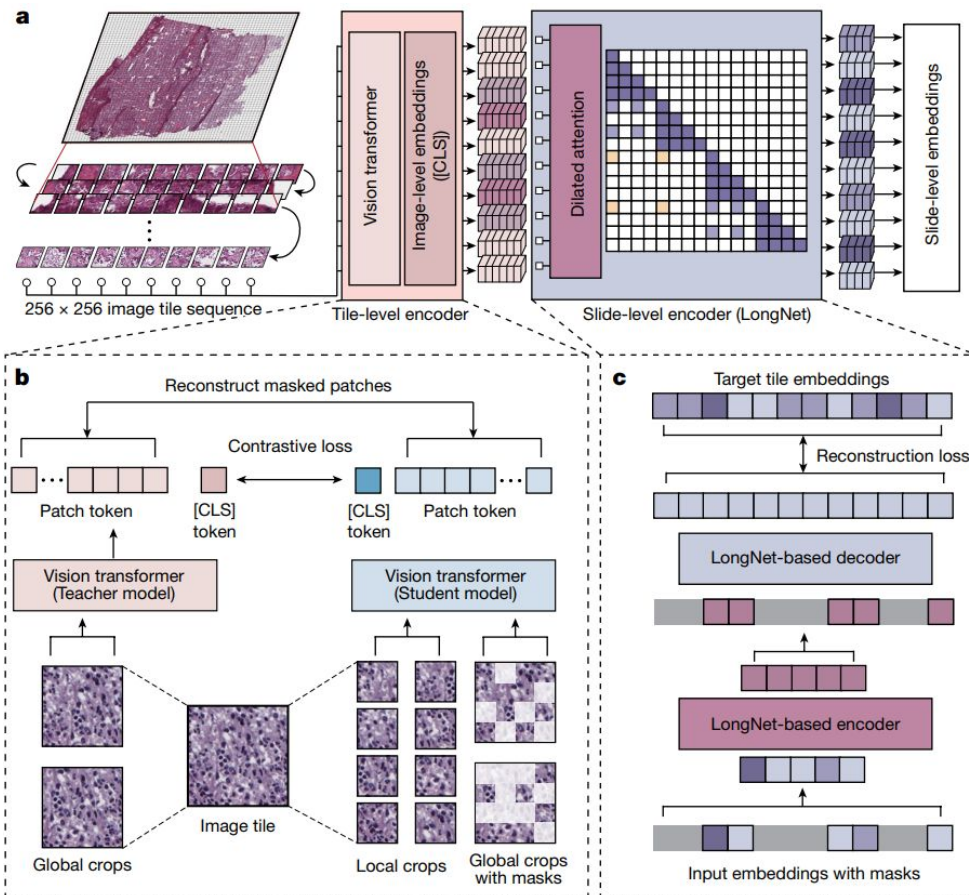
Source: Lu, A multimodal generative AI copilot for human pathology, 2024 (Harvard)

# Multimodal Tile Foundation Models

Model Name	Additional Modality	Pairs
PLIP*	Text descriptions	208k
CONCH*	Image captions	1.17m
PathChat	Image captions + Q&A	1.18m
MUSK*	Clinical reports	1m

\* open weights

# Slide-Level Foundation Models



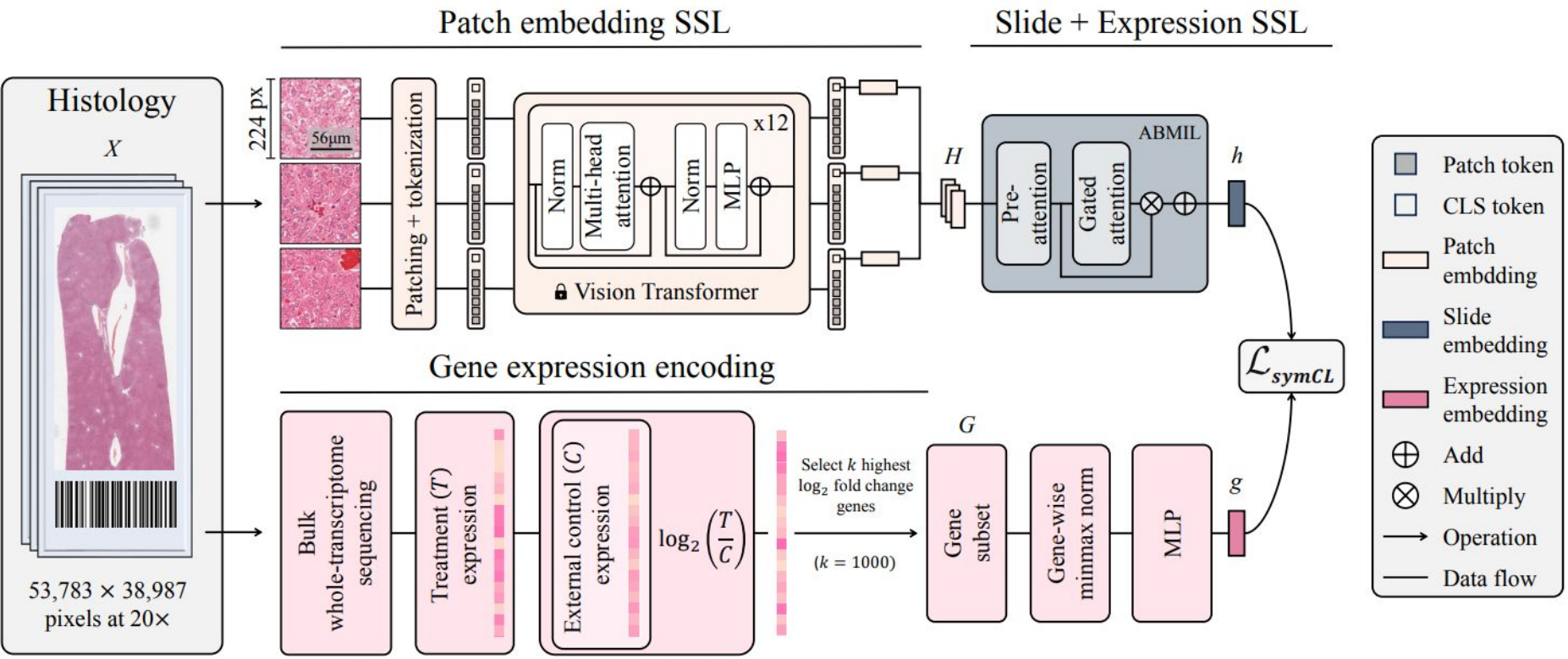
## Prov-GigaPath:

- Tile-encoder trained with DINOv2
- Slide-encoder trained with masked autoencoder
- LongNet to handle thousands of tiles per slide
- Trained on 171k WSIs from 30k patients

Source: Xu, A whole-slide foundation model for digital pathology from real-world data, 2024 (Microsoft)

# Slide-Level Multimodal: H&E + Gene Expression

TANGLE



Source: Jaume, Transcriptomics-guided Slide Representation Learning in Computational Pathology, 2024 (Harvard)

# Slide-Level Models

Model Name	Additional Modality	SSL Algorithm	WSIs
GigaSSL*	Clinical reports	SimCLR	12k
PRISM*		Contrastive	587k
Prov-GigaPath*		Masked autoencoder	171k
MADELEINE*	IHC	Contrastive	16k
TANGLE*	Gene expression	Contrastive	9k
mSTAR	Path reports, RNASeq	Contrastive	26k
CHIEF*	Multiple FM embeddings	?	61k
COBRA*		MoCo-V3	3k
TITAN*		iBOT	335k

\* open weights

# Summary

Why use foundation models for pathology?

- Weakly supervised whole slide images
- Scarce labeled data
- Robustness to distribution shifts
- Rapid prototyping

How?

- Linear probing
- Fine-tune
- Additional pre-training
- Pre-train from scratch

Going beyond tiles:

- Multimodal
- Slide-level



Ready to get started?

# How I work with clients

Who I work with:

- Founders and other leaders
- Their technical team

Example results:

- Streamline model development
- Keep up with AI trends and innovations
- Boost investor confidence

Advisory services:

- Monthly strategy call
- Weekly office hours
- Private Slack channel
- And more

# Bonus Offer

## **Team Workshop:** Harnessing the Power of Foundation Models for Pathology

In just 90 minutes, you'll gain:

- Deep insights into cutting-edge foundation model architectures
- Practical strategies for using foundation models in your workflow
- A concrete action plan for starting with foundation models

\$1,500 (25% discount if booked in the next 48 hours)

# Resources

Team workshop bonus offer (25% discount if booked in the next 48 hours):

<http://pixelscientia.com/workshops/pathology-foundation-models>

Other consulting services: [heather@pixelscientia.com](mailto:heather@pixelscientia.com)