# StainNet: A Special Staining Self-Supervised Vision Transformer for Computational Pathology

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#### Abstract

Foundation models trained with self-supervised learning (SSL) on large-scale histological images have significantly accelerated the development of computational pathology. These models can serve as backbones for region-of-interest (ROI) image analysis or patch-level feature extractors in whole-slide images (WSIs) based on multiple instance learning (MIL). Existing pathology foundation models (PFMs) are typically pre-trained on Hematoxylin-Eosin (H&E) stained pathology images. However, images with special stains, such as immunohistochemistry, are also frequently used in clinical practice. PFMs pre-trained mainly on H&E-stained images may be limited in clinical applications involving special stains. To address this issue, we propose StainNet, a specialized foundation model for special stains based on the vision transformer (ViT) architecture. StainNet adopts a selfdistillation SSL approach and is trained on over 1.4 million patch images cropping from 20,231 publicly available special staining WSIs in the HISTAI database. To evaluate Stain-Net, we conduct experiments on an in-house slide-level liver malignancy classification task and two public ROI-level datasets to demonstrate its strong ability. We also perform few-ratio learning and retrieval evaluations, and compare StainNet with recently larger PFMs to further highlight its strengths. We have released the StainNet model weights at: https://huggingface.co/JWonderLand/StainNet.

**Keywords:** Special Staining, foundation model, computational pathology.

## 1. Introduction

Modern computational pathology involves the use of deep learning models, such as convolutional neural networks (He et al., 2016) and ViT (Dosovitskiy, 2020), for clinical tasks on

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histopathological images, including cancer screening (Bejnordi et al., 2017), tumor analysis (Lu et al., 2021), morphological retrieval (Chen et al., 2022a), and survival prognosis (Chen et al., 2021a). In recent years, SSL methods have gained attention and have been proven to be highly effective (Kang et al., 2023). These methods design pretext tasks, such as contrastive learning (Chen et al., 2020) and image reconstruction (He et al., 2022), allowing the model to learn effective representations from unlabeled data and obtain good initialization weights, which can then be directly used as foundation models for pathology image feature extraction.

Most existing PFMs are pre-trained on H&E-stained histological images, which are routinely used and easily accessible in clinical workflows. As a result, these PFMs typically perform well with H&E-stained images. For example, by directly analyzing expert-annotated H&E-stained ROIs, PFMs can be used to identify tumor malignancy or microenvironment patterns (Lin et al., 2025). Additionally, when H&E-stained resection or biopsy WSI are cropped into patches, PFMs can extract these into high-dimensional features, which are used in weakly-supervised learning tasks based on MIL (Xu et al., 2025), such as cancer screening or biomarker detection (Campanella et al., 2025). However, there are other staining techniques in clinical practice. For example, by using antibodies that bind to proteins in tissue cells, immunohistochemistry (IHC) slides are widely used to visualize the presence, location, and expression levels of specific proteins, which are commonly utilized for precision cancer diagnosis, disease subtyping, and drug selection. Due to significant differences in nuclear color, tissue patterns, and texture compared to H&E staining, directly transfering existing PFMs to special-stained images for representation may be limited in clinical performance (Li et al., 2024b).

To address this issue, we propose StainNet, a ViT-based PFM specifically pre-trained on special-staining images. Specifically, to train StainNet, we first select 20,231 non-H&E-stained WSIs from the publicly available large-scale HISTAI database (Nechaev et al., 2025). For each WSI, we randomly crop up to 100 tissue-containing patch images to create our pre-training dataset. Next, to develop StainNet, we utilize DINO (Caron et al., 2021), a lightweight self-supervised learning framework specifically designed for ViT models, enabling StainNet to learn task-agnostic morphological information from a large collection of unlabeled special staining images. Finally, to evaluate StainNet, we construct downstream task datasets at both the ROI and WSI levels and compare its performance with existing pathology foundation models. Through this work, we aim to explore the adaptability of PFMs in clinical tasks involving special-stained images and call for the broader exploration of computational pathology in more diverse clinical pathology image datasets.

# 2. Related works

#### 2.1. Self-supervised PFMs

Foundation models are accelerating the advancement of computational pathology. By performing SSL on large-scale unlabeled pathology images, models can obtain histology-specific weights, which significantly improve their performance on downstream pathology tasks. For instance, CTransPath (Wang et al., 2022) uses a MoCov3-based SSL approach (Chen et al., 2021b) and is trained on 15.6 million image patches from 25 anatomical sites to demonstrate its ability in H&E staining image retrieval. HIPT (Chen et al., 2022b) and Lunit (Kang

et al., 2023) adopt DINO (Caron et al., 2021), validating the feasibility of pre-training ViT models on histological images. Given the easy scalability of ViT parameters, recent works have aimed to train on larger datasets with larger ViT models. Examples include UNI (Chen et al., 2024), Virchow (Vorontsov et al., 2024), and Prov-Gigapath (Xu et al., 2024), which aim to diversify the pre-training dataset with more varied pathology images and leverage more computational resources to build stronger general pathology representation models. Additionally, some studies explore SSL in conjunction with special staining, such as PathoDuet (Hua et al., 2024), which utilizes registered H&E and IHC images to help the model learn cross-staining features, aiding the transfer to more advanced nuclear information.

#### 2.2. Special staining in clinical pathology

H&E staining is the most commonly used histological staining method, providing basic structural information by staining the cell nuclei and cytoplasm. However, H&E staining does not fully reveal certain cell features or tissue details, especially in the detection of immune responses or protein expression. As a result, several special staining methods have been developed to address these limitations (Gridley, 1957). Common special staining methods, such as IHC, Masson's trichrome, and acid phosphatase staining, can highlight specific proteins, cell subtypes, or extracellular matrix components. For example, IHC utilizes antibodies to bind to target proteins, making it a valuable tool for cancer diagnosis and disease subtyping (Magaki et al., 2018); Masson's trichrome staining is employed to visualize elastic fibers in tissues, particularly in fibrosis-related research (Lefkowitch, 2006). Recently, the development of computational pathology has accelerated research into the application of computer vision models to these special staining images, such as lymphocyte detection in IHC images (Swiderska-Chadaj et al., 2019) and the use of multiple special stains for kidney biopsy evaluation (Jayapandian et al., 2021). Some studies have also explored the direct use of generative models to convert H&E images into special stains to improve diagnostic accuracy (De Haan et al., 2021; Bai et al., 2023; Pati et al., 2024; Yan et al., 2023). However, these special staining images differ significantly from H&E staining in terms of staining mechanisms, texture features, and visual appearance, making it difficult for large-scale pre-trained models based on general H&E staining images to capture the specific features of special staining images (Li et al., 2024b).

# 3. Methodology

We develop and evaluate our StainNet through data collection, model pre-training, and downstream testing. The entire process is illustrated in Figure 1.

#### 3.1. Data collection

We use the publicly available large-scale WSI database HISTAI (Nechaev et al., 2025) as our pre-training data. The original HISTAI dataset contains 112,801 WSIs, of which 20,265 are special stains. We first apply the OTSU thresholding algorithm (Otsu et al., 1975) to exclude 34 special staining WSIs that can not extract foreground tissue, resulting in 20,231 WSIs. Then, for each WSI, we perform non-overlapping  $224 \times 224$  pixel sliding

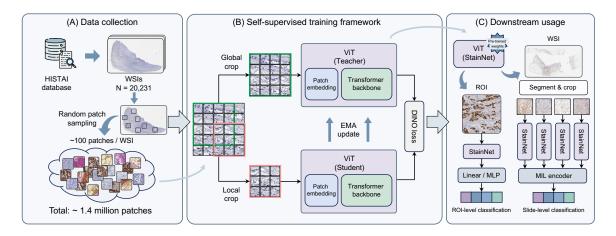


Figure 1: Overview of our proposed StainNet. We first collect approximately 1.4 million patch images with a resolution of  $224 \times 224$  pixels from the HISTAI database. Then, we train StainNet using the DINO SSL method. Finally, we adapt StainNet to downstream ROI and WSI tasks to explore its performance.

window operations on the foreground tissue, and randomly crop 100 image patches (if a WSI contains fewer than 100 patches, all the patches will be selected). In total, we obtain 1,418,938 patches for subsequent StainNet pre-training.

## 3.2. Model pre-training

Our proposed StainNet adopts the SSL method DINO (Caron et al., 2021) as the pretraining strategy to let the ViT model learn robust feature representations from specialstaining pathology images without manual annotations. DINO consists of two networks with the same architecture: a student network optimized via gradient descent and a teacher network updated using an exponential moving average (EMA) of the student weights. We follow the original DINO approach for data augmentation and multi-crop strategies. Specifically, we generate a set of global and local views for each pathology image, and the student takes all views as input, while the teacher only receives the global views. By minimizing the cross-entropy loss between the output probability distributions of the two networks, the student is encouraged to infer global tissue context solely from local information. In addition, centering and sharpening operations are applied to the teacher outputs to prevent mode collapse. Figure 2 illustrates some examples of original images and their augmented versions. The detailed hyperparameters are provided in the implementation.

#### 3.3. Downstream adaptation

Our pre-trained StainNet can be applied to both ROI and WSI analysis for special staining images. For ROI tasks, the extracted features from StainNet can be passed to either a single linear layer (linear probe) or a multilayer perceptron (MLP probe) consisting of two linear layers with a  $ReLU(\cdot)$  activation function for downstream classification. For WSI tasks, we first crop a set of tissue-containing patch images and then perform offline feature extraction

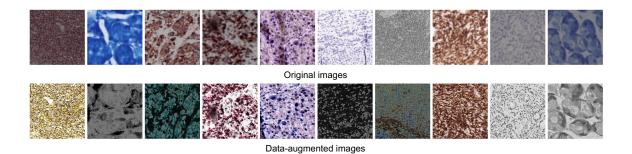


Figure 2: A set of original special staining images and their data-augmented versions.

using StainNet. The resulting  $N \times C$  feature sequence (where N is the number of image patches and C is the feature dimension) can be fed into a series of MIL models to obtain slide-level representations and classification scores.

# 4. Experiments

## 4.1. Downstream datasets and implementation

We evaluate StainNet on one private WSI task NTUH-Ki67-Liver, and two publicly available ROI tasks BCI (Liu et al., 2022) and MIST (Li et al., 2023). The NTUH-Ki67-Liver 3-class dataset consists of 46 Ki67 benign liver slides, 43 Ki67 hepatocellular carcinoma slides, and 50 Ki67 normal liver slides. All slides are collected from the Affiliated Hospital of Nantong University and scanned using the SQS-120P scanning system from Shenzhen Shengqiang Technology Co., Ltd. The annotations are performed and verified at the slide level by two pathologists. The BCI dataset is a collection of 4,870 paired H&E-IHC images showing HER2 biomarker expression. We use the IHC images and categorize them into four HER2 expression levels: 0, 1+, 2+, and 3+, with 240, 1,153, 2,142, and 1,355 samples, respectively. The MIST dataset is a large-scale H&E-IHC paired image dataset. We use the IHC images corresponding to four biomarkers: 5,642 HER2, 5,361 Ki67, 5,153 ER, and 5,139 PR to construct a 4-class dataset for evaluating the ability of PFMs to identify biomarkers across different staining conditions.

We use the AdamW optimizer with a learning rate of  $5 \times 10^{-4}$  to pre-train StainNet for 100 epochs with a batch size of 256. For multi-cropping, global crops are sampled using a random resize scale in the range of 0.4-1.0, and 8 local crops are generated with a scale in the range of 0.05-0.4. For EMA design, we increase teacher momentum linearly from 0.9995 to 1.0 throughout training. The weight decay is scheduled from 0.04 to 0.4, and the teacher temperature is set to 0.04 without warm-up and last layer freezing. We use ViT-Small, pre-trained on ImageNet datasets (Deng et al., 2009), as the initial backbone and weights for our StainNet. We also enable normalization of the last layer to stabilize the training process. The experiment is conducted on a single NVIDIA RTX PRO 6000 96GB GPU with mixed-precision and takes approximately 3.04 days to complete. We select the final epoch checkpoint as the StainNet weights used in all subsequent experiments.

For downstream fine-tuning, we use AdamW with a learning rate of  $10^{-4}$  and a weight decay of  $10^{-4}$ . We set 20 epochs with a batch size of 1 for the WSI task and 20 epochs

Patch encoder	MIL encoder										
Bal acc.	ABMIL	$\operatorname{SiMLP}$	TransMIL	WiKG	AMDMIL	S4MIL	Overall				
ResNet-50	0.742 <sub>0.093</sub>	$0.712_{0.055}$	$0.829_{0.055}$	$0.836_{0.059}$	$0.838_{0.071}$	0.832 <sub>0.076</sub>	0.7980				
CTransPath	$0.668_{0.050}$	$0.757_{0.105}$	$0.858_{0.040}$	$0.860_{0.041}$	$0.802_{0.018}$	$0.819_{0.043}$	0.7940				
PathoDuet	$0.655_{0.033}$	$0.641_{0.024}$	$0.699_{0.033}$	$0.738_{0.032}$	$0.716_{0.062}$	$0.725_{0.030}$	0.6957				
HIPT	$0.706_{0.086}$	$0.753_{0.053}$	$0.814_{0.050}$	$0.815_{0.058}$	$0.835_{0.020}$	$0.843_{0.017}$	0.7943				
Lunit	0.747 <sub>0.048</sub>	$0.838_{0.041}$	$0.872_{0.051}$	$0.891_{0.044}$	$0.872_{0.020}$	$0.862_{0.032}$	0.8470				
StainNet (Ours)	0.801 <sub>0.092</sub>	$0.873_{0.042}$	$0.896_{0.073}$	$0.916_{0.030}$	$0.892_{0.051}$	$0.883_{0.060}$	0.8769				
Patch encoder	MIL encoder										
AUC	ABMIL	$\operatorname{SiMLP}$	${\bf TransMIL}$	WiKG	AMDMIL	S4MIL	Overall				
ResNet-50	0.884 <sub>0.040</sub>	0.892 <sub>0.041</sub>	0.899 <sub>0.049</sub>	$0.916_{0.022}$	$0.909_{0.024}$	0.912 <sub>0.043</sub>	0.9019				
CTransPath	$0.876_{0.028}$	$0.885_{0.041}$	$0.940_{0.031}$	$0.940_{0.033}$	$0.912_{0.029}$	$0.923_{0.031}$	0.9127				
PathoDuet	0.815 <sub>0.056</sub>	$0.848_{0.018}$	$0.856_{0.020}$	$0.838_{0.059}$	$0.821_{0.076}$	$0.852_{0.028}$	0.8382				
HIPT	$0.838_{0.113}$	$0.884_{0.035}$	$0.906_{0.042}$	$0.907_{0.012}$	$0.914_{0.014}$	$0.930_{0.038}$	0.8964				
Lunit	0.854 <sub>0.095</sub>	$0.936_{0.021}$	$0.948_{0.031}$	$0.947_{0.029}$	$0.930_{0.019}$	$0.950_{0.026}$	0.9275				
StainNet (Ours)	0.897 <sub>0.053</sub>	$0.948_{0.029}$	$0.960_{0.036}$	$0.964_{0.020}$	$0.959_{0.030}$	$0.968_{0.027}$	0.9492				
Patch encoder	MIL encoder										
F1-score	ABMIL	$\operatorname{SiMLP}$	${\bf TransMIL}$	WiKG	AMDMIL	S4MIL	Overall				
ResNet-50	0.737 <sub>0.093</sub>	$0.689_{0.059}$	$0.828_{0.052}$	0.835 <sub>0.060</sub>	0.837 <sub>0.066</sub>	0.833 <sub>0.077</sub>	0.7931				
CTransPath	$0.556_{0.084}$	$0.726_{0.122}$	$0.856_{0.042}$	$0.852_{0.045}$	$0.791_{0.026}$	$0.821_{0.040}$	0.7671				
PathoDuet	$0.604_{0.077}$	$0.612_{0.030}$	$0.660_{0.069}$	$0.716_{0.038}$	$0.691_{0.073}$	$0.718_{0.036}$	0.6668				
HIPT	$0.657_{0.126}$	$0.731_{0.072}$	$0.813_{0.049}$	$0.810_{0.065}$	$0.836_{0.019}$	$0.834_{0.027}$	0.7801				
Lunit	0.719 <sub>0.095</sub>	$0.838_{0.043}$	$0.870_{0.054}$	$0.893_{0.044}$	$0.867_{0.020}$	$0.860_{0.028}$	0.8409				
StainNet (Ours)	0.777 <sub>0.141</sub>	$0.870_{0.038}$	$0.891_{0.076}$	$0.914_{0.030}$	$0.890_{0.058}$	<b>0.881</b> <sub>0.061</sub>	0.8706				

Table 1: Comparison results of our StainNet with five similarly sized PFMs on the NTUH-Ki67-Liver classification dataset. We use six different MIL methods for slide-level fine-tuning and report average balanced accuracy with standard deviation across all folds.

with a batch size of 128 for the ROI task. Early stopping is set to 5 epochs for both functions. For classification tasks, we save the fine-tuned weights and report accuracy, balanced accuracy, AUC, and F1-score from each epoch that achieves the best balanced accuracy. For retrieval tasks, we use cosine similarity for image matching and report recall@K and mAP@K, where K ranges from 1 to 20. All experiments are conducted using stratified 5-fold cross-validation and full-precision computing on a single NVIDIA RTX 3090 24GB GPU. All baseline methods are trained using the same settings.

# 4.2. Comparison with PFMs of similar parameter scale

We compare the proposed StainNet with five PFMs of comparable parameter size: CTransPath (Wang et al., 2022), PathoDuet (Hua et al., 2024), HIPT (Chen et al., 2022b), Lunit (Kang

Patch encoder	Linear probe				MLP probe			
$BCI\ 4\ classes\ datasets$	Acc.	$Bal \ \overline{acc.}$	$\overline{AUC}$	F1-score	Acc.	$Bal\ \overline{acc.}$	$\overline{AUC}$	F1-score
ResNet-50	0.587 <sub>0.010</sub>	0.407 <sub>0.010</sub>	0.774 <sub>0.018</sub>	0.541 <sub>0.014</sub>	0.709 <sub>0.016</sub>	0.614 <sub>0.019</sub>	0.900 <sub>0.008</sub>	0.703 <sub>0.016</sub>
CTransPath	$0.575_{0.011}$	$0.387_{0.007}$	$0.808_{0.007}$	$0.499_{0.008}$	$0.736_{0.015}$	$0.653_{0.021}$	$0.913_{0.008}$	$0.733_{0.015}$
PathoDuet	0.6120.016	$0.430_{0.015}$	$0.810_{0.014}$	$0.554_{0.028}$	$0.687_{0.019}$	$0.564_{0.044}$	$0.877_{0.012}$	$0.664_{0.024}$
HIPT	$0.564_{0.023}$	$0.404_{0.023}$	$0.754_{0.020}$	$0.532_{0.028}$	$0.661_{0.009}$	$0.533_{0.010}$	$0.859_{0.010}$	$0.650_{0.011}$
Lunit	0.696 <sub>0.015</sub>	$0.545_{0.018}$	$0.880_{0.008}$	$0.681_{0.017}$	$0.872_{0.016}$	$0.816_{0.033}$	$0.976_{0.002}$	$0.871_{0.017}$
StainNet (Ours)	0.7090.025	$0.597_{0.036}$	$0.892_{0.023}$	$0.701_{0.025}$	$0.874_{0.010}$	$0.839_{0.020}$	$0.979_{0.004}$	$0.873_{0.010}$
Patch encoder	<u> </u>	Linear	probe		<u>'</u>	MLP	probe	
Patch encoder  MIST 4 classes datasets	Acc.	$\frac{\text{Linear}}{Bal\ acc.}$	probe AUC	F1-score	Acc.	$\begin{array}{c} \underline{\text{MLP}} \\ Bal \ acc. \end{array}$	probe AUC	F1-score
	Acc. 0.597 <sub>0.009</sub>			F1-score 0.593 <sub>0.008</sub>	Acc. 0.671 <sub>0.005</sub>			F1-score 0.671 <sub>0.005</sub>
MIST 4 classes datasets	l	$Bal \overline{acc.}$	AUC		l	Bal acc.	AUC	
MIST 4 classes datasets ResNet-50	0.597 <sub>0.009</sub>	Bal acc. 0.592 <sub>0.009</sub>	AUC 0.835 <sub>0.002</sub>	0.593 <sub>0.008</sub>	0.671 <sub>0.005</sub>	Bal acc. 0.667 <sub>0.004</sub>	AUC 0.884 <sub>0.002</sub>	0.671 <sub>0.005</sub>
MIST 4 classes datasets  ResNet-50 CTransPath	0.597 <sub>0.009</sub> 0.663 <sub>0.008</sub>	Bal acc.  0.592 <sub>0.009</sub> 0.658 <sub>0.008</sub>	AUC 0.835 <sub>0.002</sub> 0.878 <sub>0.004</sub>	$0.593_{0.008} \\ 0.659_{0.006}$	$ \begin{array}{ c c c c c c c c c c c c c c c c c c c$	Bal acc.  0.667 <sub>0.004</sub> 0.728 <sub>0.007</sub>	AUC 0.884 <sub>0.002</sub> 0.920 <sub>0.004</sub>	$0.671_{0.005} \\ 0.731_{0.007}$
MIST 4 classes datasets  ResNet-50  CTransPath  PathoDuet	0.597 <sub>0.009</sub> 0.663 <sub>0.008</sub> 0.590 <sub>0.010</sub>	Bal acc. 0.592 <sub>0.009</sub> 0.658 <sub>0.008</sub> 0.585 <sub>0.011</sub>	AUC 0.835 <sub>0.002</sub> 0.878 <sub>0.004</sub> 0.827 <sub>0.006</sub>	$0.593_{0.008} \\ 0.659_{0.006} \\ 0.586_{0.013}$	$ \begin{array}{c c} 0.671_{0.005} \\ 0.732_{0.007} \\ 0.664_{0.013} \end{array} $	Bal acc.  0.667 <sub>0.004</sub> 0.728 <sub>0.007</sub> 0.660 <sub>0.012</sub>	AUC 0.884 <sub>0.002</sub> 0.920 <sub>0.004</sub> 0.885 <sub>0.004</sub>	$0.671_{0.005} \\ 0.731_{0.007} \\ 0.664_{0.011}$

Table 2: Comparison results of our StainNet with five similarly sized PFMs on the BCI and MIST classification datasets. We use linear probe and MLP probe methods for ROI-level fine-tuning and report four average evaluation metrics with standard deviation across all folds.

et al., 2023), and ResNet-50 (ImageNet pre-trained) (He et al., 2016). For the WSI task, we evaluate six MIL models: (1) ABMIL (Ilse et al., 2018), a gated-attention MIL,(2) SiMLP (Li et al., 2025), a method combining unsupervised average pooling with an MLP classifier, (3) TransMIL (Shao et al., 2021), a transformer-based MIL, (4) WiKG (Li et al., 2024a), a dynamic graph representation method, (5) AMDMIL (Ling et al., 2024), a method combining masked denosing with agent tokens, and (6) S4MIL (Fillioux et al., 2023), a state space model-based MIL. For the ROI tasks, we freeze each PFM and apply two probe settings: a linear probe and an MLP probe for fine-tuning. Table 1 and Table 2 respectively show the results on the slide-level NTUH-Ki67-Liver task and ROI-level BCI and MIST tasks.

For NTUH-Ki67-Liver, StainNet consistently achieves the best overall performance across all MIL encoders. We also observe that lightweight MIL encoders exacerbate the performance gap between H&E-stained image pre-trained PFMs, whereas more complex MIL encoders mitigate this gap. For example, StainNet outperforms the second-best PFM in terms of balanced accuracy by 5.4% with ABMIL, whereas the margin narrows to 2.5% when using WiKG. For BCI and MIST, StainNet continues to outperform significantly, especially on MIST, where StainNet surpasses the popular CTransPath by 11.7% and 15.4% in balanced accuracy under the linear and MLP probe settings.

## 4.3. Comparison with larger PFMs

We further compare StainNet with three recently proposed larger PFMs: PathOrchestra (Yan et al., 2025), GPFM (Ma et al., 2025), and UNI (Chen et al., 2024) as shown in Figure 3. Although StainNet has nearly 14× fewer parameters than them (22M vs 303M), it still

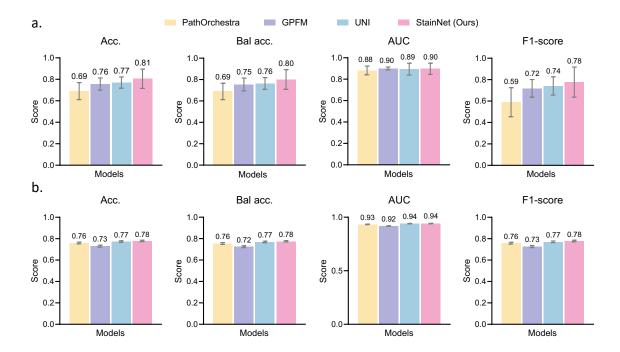


Figure 3: Comparison of StainNet with three larger PFMs across four evaluation metrics: accuracy, balanced accuracy, AUC, and F1-score. **a.** Performance on the NTUH-Ki67-Liver slide-level classification task with ABMIL fine-tuning. **b.** Performance on the MIST ROI-level classification task with linear probe fine-tuning.

achieves the best performance. For example, StainNet surpasses these three large PFMs by 3.72%, 4.70%, and 11.15% in terms of balanced accuracy (80.09% vs. 76.37%, 75.39%, and 68.94%). These results demonstrate that representations learned from large-scale special-stain images provide substantial advantages over H&E-based PFMs when applied to special-stain tasks.

## 4.4. Comparison of image retrieval

We also examine the retrieval capability of StainNet to assess its semantic representation quality on special staining histology images. We compare StainNet with CTransPath and UNI on the MIST dataset, as shown in Figure 4. We observe that StainNet consistently achieves the best Recall@K and mAP@K across all choices of K, indicating that it has learned a well-structured embedding space for special staining images.

## 4.5. Effectiveness of different fine-tuning data ratios

We further evaluate the fine-tuning performance of StainNet under varying amounts of downstream special-stain training data to assess its learning capability. Figure 5 presents the results of StainNet, CTransPath, and UNI on the MIST dataset using linear and MLP probes under different proportions of fine-tuning samples. We observe that although Stain-

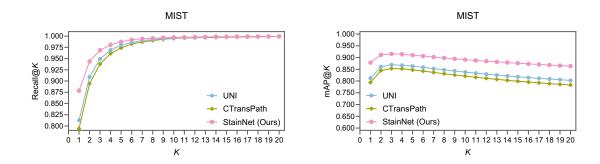


Figure 4: Comparison of image retrieval performance among CTransPath, UNI, and our StainNet on the MIST ROI-level dataset.

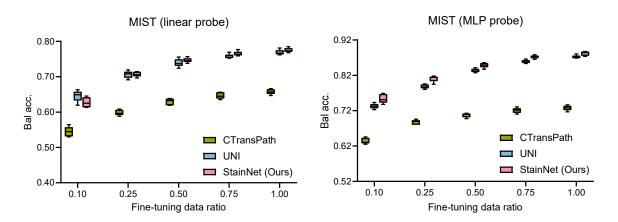


Figure 5: Linear probe and MIL probe fine-tuning results of CTransPath, UNI and our StainNet on the MIST ROI-level dataset across varying training data ratios.

Net performs slightly worse than UNI under the 10% data setting with a linear probe, it achieves the best performance in all other settings and substantially outperforms the CTransPath of similar parameter size. Notably, applying an MLP probe significantly increases the performance gap between StainNet and UNI in low-fine-tuning data scenarios.

#### 4.6. Performance variation under different pre-training iterations

Finally, we investigate the performance variation of StainNet under different pre-training iterations and the behavior of its loss curve. We believe these observations can provide valuable insights for the development of SSL methods in computational pathology, where pre-training details such as the expected loss curve and convergence behavior are often underreported. Figure 6 illustrates the pre-training loss curve over 100 epochs and the corresponding linear probe performance on the MIST dataset at six iteration checkpoints. We observe that the loss plateaus between epoch 30 and 50, but with significant performance improvement during this period. After epoch 60, the overall loss begins to decrease again, but with slight changes in performance. Ultimately, the model reaches its optimal state

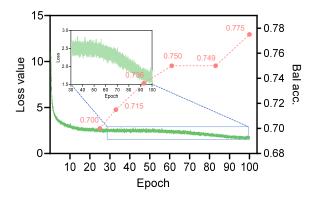


Figure 6: Pre-training loss curve of StainNet and comparison of linear probe performance on MIST under six different iterations.

at the final iteration. These findings suggest that the shift in the DINO pre-training loss may serve as a valuable indicator of representation quality. While additional exploration is required to generalize these observations to other settings, we encourage future SSL development in computational pathology to consider allocating more training epochs, which may be crucial for achieving optimal model performance.

#### 5. Conclusion and future works

In this work, we propose StainNet, a lightweight pathology foundation model specifically for special staining histopathology image representation. Utilizing the DINO self-supervised framework, StainNet is trained from over 1.4 million unlabeled images. We evaluate StainNet on one in-house WSI task and two public ROI-level tasks, including classification, few-ratio settings, and image retrieval. The results demonstrate that StainNet learns strong and robust representations for special staining pathology images.

For future work, we plan to develop a larger StainNet by expanding the pre-training dataset, scaling the model parameter size, and evaluating it on broader downstream tasks. We will also explore integrating StainNet with H&E-based PFMs for multimodal applications such as survival prognosis and therapy response prediction. We believe that StainNet will further accelerate the research and applications in computational pathology involving diverse histological modalities.

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