# **1** Supplementary Material

## 1.1 Multimodal Pre-training Methods

Medical vision-language pre-training enhances medical image analysis by learning domain-specific features from medical images paired with clinical descriptions. By jointly encoding images and reports, these models better understand visual and textual information, improving performance and interpretability. Typical methods improve image-text contrastive learning **3[7]18[34]**, align image and text embeddings using semantic labels **[31]**, or enhance image representation through masked image and language modeling **[36]**. Recent methods have focused on radiology, especially chest X-rays **[22]32[33]**, due to the abundance of image-report pairs that help learn the relationship between visual features and medical findings. However, this approach is less applicable in other medical domains like ophthalmology, where retinal images have diverse modalities and generally lack accompanying text information.

Unlike RETFound [37] and FLAIR [27], we propose a universal retinal FM that processes multiple imaging modalities and integrates various expert annotations into the image encoder. By leveraging multimodal images and domain knowledge, this model enables comprehensive representations, facilitates multimodal reasoning, captures broader anatomical and physiological relationships, and reduces development and maintenance costs.

#### 1.2 Dataset Preparation

**Pre-training Dataset.** Based on FLAIR, we collected a large dataset (Tabel. comprising 187,270 publicly accessible CFP and OCT images for the pre-training of our foundation model and the experiments conducted. More details can be found in FLAIR 27.

**Fine-tuning Dataset.** To conduct a comprehensive evaluation of the foundation model, we collected 7 CFP datasets and 1 OCT dataset according to the experimental setup defined by RETFound, and divided them following the data division ratios provided by RETFound <u>37</u>.

Task Specific Dataset. Based on the labels in the pre-training dataset, we constructed a task-specific dataset for Diabetic Retinopathy classification, which includes images from EYEPACS, PARAGUAY, OIA-DDR, and Deep-DRiD, totaling 51,556 images. Similarly, a task-specific dataset for OCT disease classification was developed based on the OCTCELL dataset.

## 1.3 Expert Knowledge Descriptions

For the domain knowledge descriptors related to retinal diseases based on CFP, we referred to FLAIR [27] for guidance. Meanwhile, for the domain knowledge descriptors concerning retinal diseases based on OCT, we utilized ChatGPT-4 to summarize four distinct descriptions for the corresponding disease label names, which were then employed as the domain knowledge descriptors (Tabel. [2]).

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Table 1: Collected publicly available dataset for foundation model pre-training.

No.	Name	$\mathbf{Count}$	Labels
1	OCTCELL 16	83,484	CNV, DME, DRUSEN, and NORMAL
2	EYEPACS 11	35,126	noDR, mildDR, modDR, sevDR, prolDR
3	RFMid 25	3,170	DR, ARMD, MH, DN, MYA, BRVO, TSLN, ERM, LS, MS CSR,
			ODC, CRVO, TV, AH, ODP, ODE, ST, AION, PT, RT RS, CRS,
			EX, RPEC, MHL, RP, CWS, CB, ODM, PRH, MNF, HR, CRAO,
			TD, CME, PTCR, CF, VH, MCA VS, BRAO, PLQ, HPED, CL
4	EYENET 15	15,709	Text
5	LAG 19	4,854	G, noG
6	ODIR 1	10,846	N, DR, G, CAT, ARMD, HR, MYA
7	PARAGUAY 4	757	noDR, mildDR, modDR, sevDR, prolDR
8	STARE 14	397	Text
9	ARIA 12	143	N, ARMD, DR
10	AGAR300 9	28	DR, MA
11	FUND-OCT 13	179	G, N, CME, neovARMD, geoARMD, acCSR, chCSR
12	DRIONS-DB 6	110	noCAT, Dis
13	Drishti-GS1 28	101	N, G
14	E-ophta	265	EX, MA
15	G1020 2	1,020	G, N
16	HRF 5	45	N, G, DR, noisy
17	DRIGA 35	650	G, noG
18	RUC 24	10.072	MA
19	CIA-DDR 20	13,073	noDR, mildDR, modDR, sevDR, proiDR, HE, nEA, SEA, MA
20		1,219	noDR, mildDR, modDR, sevDR, proiDR, nE, nEA, SEA
21	CUAKSU 17	9,939	C $poC$
22	DB1 2 26	1 460	N ROSD DEX DN CWS SUDHE doopHE
20	ScarDat 30	007	IS noLS
25	ACRIMA 10	705	G noG
26	DeepDBiD 23	2,000	noDB mildDB modDB sevDB prolDB
	DeepDitid 20	2,000	
	Total	187,270	

# 1.4 Statistical Significance Analysis

Fig. 1 shows the statistically significant analysis of UrFound compared to the second-best results in Table 1 of the paper, based on a t-test with a p-value of 0.05. UrFound performs similarly to the second-best method on IDRID and JSIEC, and significantly better on the other six datasets.

## 1.5 External Validation

We conducted external evaluations on the IDRID, APTOS, and Messidor datasets and found that our UrFound model demonstrates strong generalizability, and outperforms RETFound and FLAIR in most cases, with statistical significance based on a t-test with a p-value of 0.05 (Fig. 2).

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Table 2: Expert	Knowledge	descriptions i	for (	OCT-based	retinal	diseases

Category	Domain Knowledge descriptor
CNV	<ol> <li>"The OCT image reveals a network of new blood vessels beneath the retinal pigment epithelium, indicative of Choroidal Neovascularization. These vessels are irregular and often associated with age-related macular degeneration."</li> <li>"There is noticeable distortion and elevation of the overlying retinal layers, which is characteristic of the leakage and bleeding from these abnormal vessels."</li> <li>"Pockets of fluid accumulation under the retina, known as subretinal fluid, are evident, causing a dome-shaped elevation of the retina."</li> <li>"Areas of hemorrhage and exudation are visible between the retinal layers and beneath the retinal pigment epithelium, indicating active vascular leakage."</li> </ol>
DME	<ol> <li>"In Diabetic Macular Edema, the OCT scan shows a significant thickening of the macula, particularly in the inner retinal layers, due to fluid accumulation. This condition is a common complication of diabetic retinopathy."</li> <li>"Multiple cystic spaces within the retinal layers are observed, filled with fluid, giving a sponge-like appearance to the retina."</li> <li>"Hyperreflective foci are seen below the retinal pigment epithelium, representing hard exudates, which are residues of lipid deposits from leaking blood vessels."</li> <li>"In advanced cases, disruption and irregularity of the retinal pigment epithelium layer are noted, likely due to chronic edema and vascular leakage."</li> </ol>
DRUSEN	<ol> <li>"Drusen appear as small, round elevations beneath the retinal pigment epithe- lium layer in OCT images. These are accumulations of extracellular material, com- monly associated with age-related macular degeneration."</li> <li>"The drusen vary in size and confluence, with larger and more numerous drusen indicating a higher risk of progression to advanced macular degeneration."</li> <li>"In cases of extensive drusen, there is noticeable distortion and thickening of the overlying retinal pigment epithelium layer."</li> <li>"Some drusen exhibit a central hyperreflective core with a surrounding hypore- flective halo, suggesting varying stages of drusen evolution."</li> </ol>
NORMAL	<ul> <li>1. "The normal retina in OCT imaging presents a well-defined, multi-layered structure. Each layer exhibits its characteristic reflectivity, with clear demarcation between layers."</li> <li>2. "The retinal pigment epithelium layer appears as a uniform, thin band adjacent to the highly reflective Bruch's membrane."</li> <li>3. "The photoreceptor layer, including the cones and rods, is orderly and shows no signs of fluid accumulation or structural distortion."</li> <li>4. "The nerve fiber layer, ganglion cell layer, and inner and outer nuclear layers all display normal thickness and reflectivity, with no signs of pathology or abnormal-lity."</li> </ul>

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Fig. 1: Analysis of Statistical Significance with the Second-Best Results in Table 1 of the Paper.



Fig. 2: Performance of RETFound, UrFound, and FLAIR on External Validation with Statistical Analysis

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