



OPEN A dual-stream deep learning framework for skin cancer classification using histopathological-inherited and vision-based feature extraction

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Skin cancer, particularly melanoma, remains one of the most life-threatening forms of cancer worldwide, with early detection being critical for improving patient outcomes. Traditional diagnostic methods, such as dermoscopy and histopathology, are often limited by subjectivity, interobserver variability, and resource constraints. To address these challenges, this study proposes a dual-stream deep learning framework that combines histopathological-inherited and vision-based feature extraction for accurate and efficient skin lesion diagnosis. The framework uses the U-Net architecture for precise lesion segmentation, followed by a dual-stream approach: the first stream employs Virchow2, a pretrained model, to extract high-level histopathological embeddings, whereas the second stream uses Nomic, a vision-based model, to capture spatial and contextual information. The extracted features are fused and integrated to create a comprehensive representation of the lesion, which is then classified via a multilayer perceptron (MLP). The proposed approach is evaluated on the HAM10000 dataset, achieving a mean accuracy of 96.25% and a mean F1 score of 93.79% across 10 trials. Ablation studies demonstrate the importance of both feature streams, with the removal of either stream resulting in significant performance degradation. Comparative analysis with existing studies highlights the superiority of the proposed framework, which outperforms traditional single-modality approaches. The results underscore the potential of the dual-stream framework to enhance skin cancer diagnosis, offering a robust, interpretable, and scalable solution for clinical applications.

Keywords Classification, Deep learning (DL), Feature extraction, Histopathology

Skin cancer is one of the most prevalent and life-threatening types of cancer worldwide, with melanoma representing its deadliest form¹. Melanoma accounts for the majority of skin cancer-related deaths, despite constituting a smaller proportion of skin cancer cases than nonmelanoma types such as basal cell carcinoma and squamous cell carcinoma^{2,3}.

According to the World Health Organization (WHO), approximately 132,000 new cases of melanoma are diagnosed annually worldwide^{4,5}, and this number continues to rise due to factors such as increased ultraviolet (UV) radiation exposure, aging populations, and improved diagnostic capabilities⁶. The aggressive nature of melanoma, coupled with its ability to metastasize rapidly, underscores the importance of early detection⁷.

When diagnosed at an early stage, the five-year survival rate for patients with melanoma exceeds 99%, but this rate decreases to less than 30% for patients with advanced-stage disease. These statistics highlight the critical need for accurate and timely diagnostic tools to improve patient outcomes and reduce mortality rates^{8,9}.

Traditional diagnostic methods for skin cancer rely on dermatologists using tools like dermoscopy and visual inspection¹⁰. Dermoscopy, which magnifies and illuminates the skin, improves accuracy but depends heavily on clinician expertise^{11,12}. Histopathological examination remains the “gold standard”, providing definitive diagnostic details through biopsy and microscopic analysis¹³. However, it is invasive, time-consuming, and inaccessible in resource-limited settings^{14,15}.

Challenges with traditional methods include interobserver variability, leading to misdiagnoses, and subjective reliance on visual interpretation^{16–18}. Histopathology can also be error-prone with inadequate samples. These

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limitations, compounded by a shortage of specialists, highlight the need for objective, scalable, and efficient diagnostic tools^{19,20}.

Whole slide imaging (WSI) has transformed pathology by digitizing glass slides, enabling applications in diagnostics, education, and research²¹. WSI facilitates telepathology and algorithm-based decision support but faces adoption challenges, including technical issues and diagnostic difficulties^{22,23}. AI integration has expanded WSI's potential, particularly in cancer diagnosis, where AI algorithms improve accuracy and reduce turnaround times^{24,25}. AI also enhances radiology, pharmacology, and personalized medicine, identifying patterns imperceptible to humans^{24,25}.

Despite its promise, skepticism about AI persists due to interpretability concerns, ethical considerations, and unclear reimbursement opportunities²³. Effective communication is needed to build trust and demonstrate AI's benefits²¹.

AI, particularly deep learning (DL), offers solutions for skin cancer diagnosis by analyzing dermatoscopic and histopathological images²⁶. Using datasets like HAM10000, AI models identify lesion patterns and assist clinicians, improving efficiency and accessibility in underserved areas^{27–29}. This integration represents a paradigm shift, enhancing outcomes^{30,31}. DL automates feature extraction, with CNNs achieving state-of-the-art performance in lesion segmentation and classification^{32–35}. For example, U-Net captures fine-grained lesion details critical for accurate classification^{36,37}. Challenges remain, such as overfitting, driven by limited, diverse datasets^{38,39}.

To address these challenges, researchers have begun exploring hybrid approaches that combine multiple DL techniques⁴⁰. For example, pretrained models such as Virchow2, which utilize histopathological data, can extract high-level embeddings that capture intricate patterns in skin lesions. Similarly, vision-based approaches such as Nomic can complement these embeddings by providing additional contextual information.

Therefore, the current study aims to address the challenges associated with skin lesion diagnosis by using advanced DL techniques and a dual-stream methodology to improve skin cancer detection accuracy, efficiency, and interpretability. Using the HAM10000 (HAM10K) dataset, this research employs the U-Net architecture for precise lesion segmentation, ensuring accurate localization of affected areas, critical for subsequent feature extraction and classification.

To enhance diagnostic performance, this study introduces a dual-stream approach: the first track uses Virchow2, a pretrained model that uses dermatoscopic data to extract high-level embeddings capturing intricate patterns in skin lesions, whereas the second track employs a vision-based approach in which Nomic is used to extract spatial and contextual information from these images. This study aims to create a comprehensive diagnostic system to improve classification accuracy and robustness by integrating these methodologies.

The rest of the manuscript is organized in the following manner: Section 2 provides an overview of the relevant research. Section 3 introduces the proposed approach for skin lesion diagnosis. Section 4 evaluates the proposed dual-stream methodology, and its performance was assessed via a comprehensive set of metrics. Finally, Section 5 concludes the work and summarizes potential directions for further research.

Related studies

The field of skin lesion classification has undergone significant advancements in recent years, with DL techniques playing a pivotal role in improving diagnostic accuracy. In 2022, Ali et al. introduced a multiclass skin cancer classification system using EfficientNets, achieving an F1 score of 87% and a top-1 accuracy of 87.91% on the HAM10000 dataset⁴¹. Their work demonstrated the effectiveness of transfer learning and fine-tuning in handling imbalanced datasets. However, the study highlighted that increased model complexity does not always lead to better performance, suggesting the need for more efficient architectures.

Similarly, Shetty et al. proposed a machine learning and convolutional neural network (CNN) approach for skin lesion classification, achieving a training accuracy of 95.18% and a testing accuracy of 86.43% with a customized CNN model⁴². Their work emphasized the importance of data augmentation and k-fold cross-validation to address class imbalance and improve model robustness. Despite these advancements, the reliance on traditional CNNs has limited their ability to capture more complex patterns in skin lesions, and the significant gap between training accuracy and testing accuracy indicates potential overfitting.

In 2024, Adebisi et al. explored multimodal learning by combining skin lesion images with patient metadata (age, sex, and lesion location) via the ALBEF model⁴³. Their approach achieved an accuracy of 94.11% and an AUCROC of 0.9426 but with a lower recall of 90.19%, outperforming traditional DL models that rely solely on images. This study demonstrated the potential of multimodal learning to improve diagnostic accuracy, particularly in primary care settings where access to expert dermatologists is limited. However, the study was limited by the limited metadata available, suggesting that incorporating additional clinical data could further enhance performance.

Recent advancements in transformer-based models have also contributed to the field. Xin et al. introduced an improved transformer network for skin cancer classification, achieving state-of-the-art performance on the ISIC dataset⁴⁴. Their work demonstrated the potential of transformers to capture long-range dependencies in dermatoscopic images, but it also highlighted the challenges of training such models on limited medical datasets.

In addition, Mao et al. proposed the medical supervised masked autoencoder (MSMAE), which introduced a better masking strategy and fine-tuning schedule for medical image classification⁴⁵. Their approach improved the efficiency of transformer-based models, making them more suitable for skin lesion classification tasks.

The reviewed studies collectively highlight several limitations that the current research aims to address. First, while models such as EfficientNets and SBXception have strong performance, they often struggle with class imbalance and computational efficiency. The current study addresses this issue by employing data augmentation and a dual-stream approach that balances accuracy and computational overhead. Second, while multimodal learning, as demonstrated by Adebisi et al., has shown promise, it is limited by the availability of metadata⁴³.

Third, while transformer-based models proposed by Xin et al.⁴⁴ and Mao et al.⁴⁵ have shown potential, they require significant computational resources. The current study integrates efficient feature extraction methods and pretrained models to reduce computational complexity while maintaining high accuracy. The current research aims to create a robust and efficient diagnostic system for skin lesion classification by addressing these limitations.

Methodology

In this section, we introduce the proposed approach for skin lesion diagnosis (see Fig. 1), which uses a dual-stream framework combining histopathological-inherited and vision-based feature extraction. The methodology begins with Data Acquisition, where dermatoscopic images are collected and preprocessed to ensure high-quality input for the model. These images are then passed through a U-Net Model for precise lesion segmentation, generating predicted regions of interest (ROIs) that highlight the affected areas.

The segmented ROIs are processed in parallel by two streams: the embedded stream network using Virchow2 for histopathologically inherited feature extraction and the vision stream network using Nomic for vision-based feature extraction. The features extracted from both streams are fused and integrated to create a comprehensive representation of the lesion, which is then used for classification. The model is trained via a carefully designed loss function and evaluated via a set of performance metrics to ensure robust and accurate diagnosis.

Preprocessing of the dataset

The HAM10K dataset, consisting of 10,015 dermatoscopic images across seven common skin lesion types, was preprocessed to ensure consistency and compatibility with the dual-stream framework. The preprocessing pipeline included several key steps to enhance data quality and facilitate robust model training.

First, all images were resized to a uniform resolution of 224×224 pixels to standardize input dimensions while preserving critical details for feature extraction. This resizing step also ensured computational efficiency during training and inference. To address class imbalance, a combination of undersampling and oversampling techniques was applied, ensuring that each skin lesion type was adequately represented in the training set.

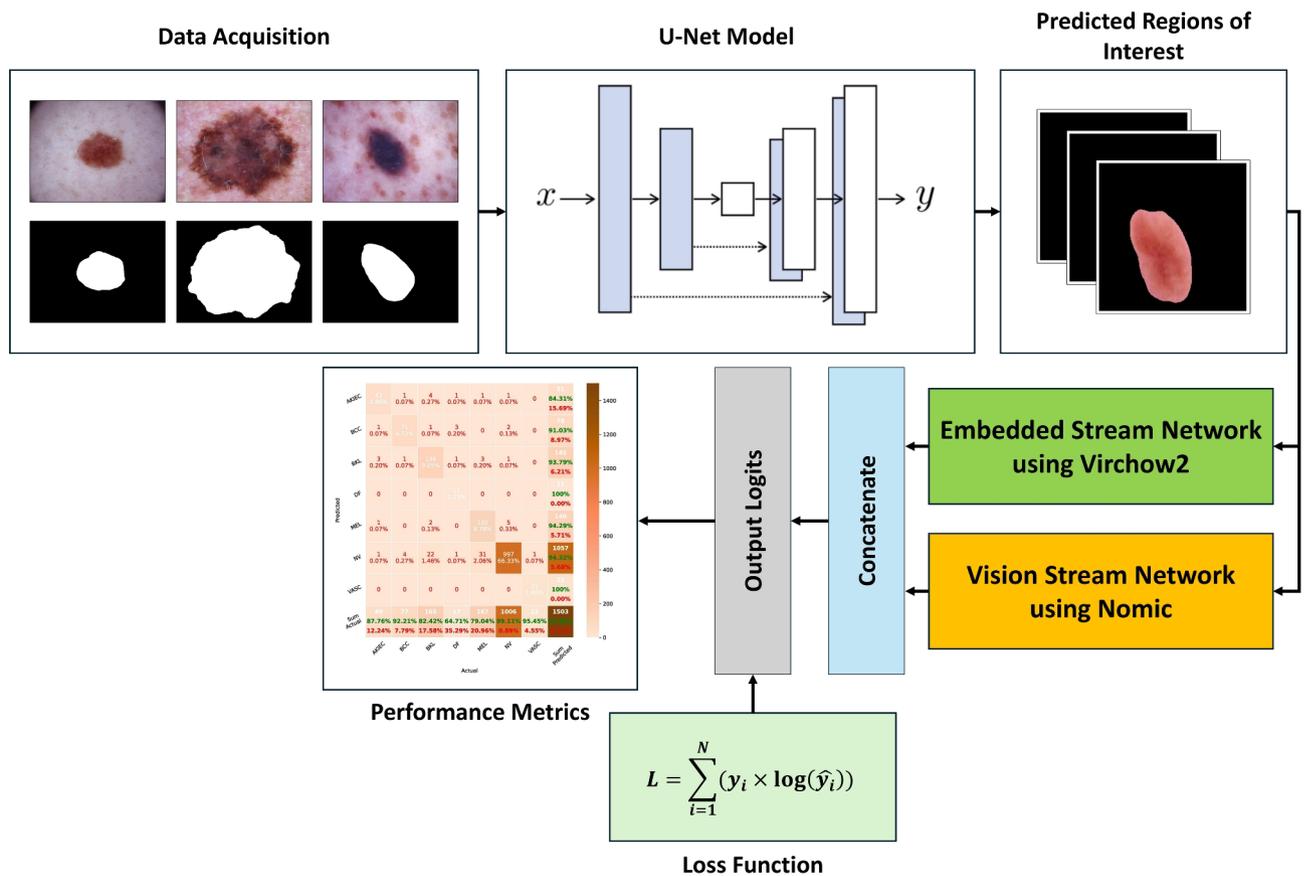


Fig. 1. Graphical representation of the proposed dual-stream framework for skin lesion diagnosis. It consists of: (1) Data Acquisition, where dermatoscopic images are collected and preprocessed; (2) U-Net Model, which performs lesion segmentation to generate predicted ROIs; (3) Embedded Stream Network, which uses Virchow2 to extract histopathologically inherited features; (4) Vision Stream Network, which uses Nomic to extract vision-based features; (5) Feature Fusion and Integration, where features from both streams are combined; and (6) Performance Evaluation, where the model’s performance is assessed via metrics such as accuracy and F1 score.

Next, data augmentation was performed to simulate real-world variations in lesion appearance and improve the model's generalizability. Augmentation techniques included random rotations, horizontal and vertical flipping, scaling, and color jittering. These transformations introduced variability in lesion orientation, size, and color, mimicking the diversity observed in clinical settings.

The dataset was then split into training, validation, and testing subsets using a stratified approach to maintain proportional representation of each lesion type across all splits. Specifically, 70% of the dataset was allocated for training, 15% for validation, and the remaining 15% for testing. This partitioning ensures a robust evaluation of the model's performance on unseen data, as described in Equation 1 where N represents the total number of images in the dataset.

$$N_{\text{train}} = 0.7 \times N, \quad N_{\text{val}} = 0.15 \times N, \quad N_{\text{test}} = 0.15 \times N \quad (1)$$

Finally, pixel intensity normalization was applied to all images to standardize the range of input values. Each image's pixel values were scaled to the range $[0, 1]$, ensuring consistent input distributions for the deep learning models. This normalization step improved convergence speed during training and enhanced the stability of the model. By implementing these preprocessing steps, the dataset was optimized for use in the dual-stream framework, enabling accurate and efficient training while ensuring the model's ability to generalize across diverse clinical scenarios.

Lesion segmentation using U-Net

Accurate lesion segmentation is a critical step in skin lesion diagnosis, as it enables precise localization of affected areas, which is essential for subsequent feature extraction and classification⁴⁶. This study employs the U-Net architecture for lesion segmentation because of its proven effectiveness in medical image analysis tasks.

U-Net is a convolutional neural network (CNN) designed explicitly for biomedical image segmentation and is characterized by its encoder-decoder structure with skip connections that facilitate the preservation of spatial information⁴⁷. The encoder captures contextual features through a series of convolutional and pooling layers. In contrast, the decoder reconstructs the spatial resolution of the feature maps to produce a pixel-wise segmentation mask⁴⁸.

The U-Net architecture is defined by the following key components: an encoder, a decoder, and a loss function. The encoder consists of a series of convolutional layers followed by max-pooling layers⁴⁹. Each convolutional layer applies a set of filters to the input image, extracting hierarchical features. The output of the i -th convolutional layer can be expressed as in Equation 2, where F_i represents the feature maps at layer i , W_i and b_i are the learnable weights and biases, $*$ denotes the convolution operation, and σ is the activation function (e.g., ReLU). The max pooling operation reduces the spatial dimensions of the feature maps, enabling the network to capture broader contextual information.

$$F_i = \sigma(W_i * F_{i-1} + b_i) \quad (2)$$

The decoder pathway reconstructs the spatial resolution of the feature maps through upsampling and convolutional layers. The upsampling operation is followed by concatenation with the corresponding feature maps from the encoder pathway via skip connections⁵⁰. This process can be described as in Equation 3, where F_j represents the feature maps at the j -th decoder layer, and $\text{Upsample}(\cdot)$ denotes the upsampling operation. The skip connections ensure that fine-grained details from the encoder are preserved, enabling precise localization of lesion boundaries.

$$F_j = \sigma(W_j * \text{Upsample}(F_{j+1}) + b_j) \quad (3)$$

The utilized U-Net model (see Fig. 2) is trained via the Dice loss function, which is particularly suitable for segmentation tasks with imbalanced class distributions⁵¹. The Dice loss is defined as in Equation 4, where y_p and \hat{y}_p represent the ground truth and predicted segmentation masks, respectively, for pixel p . Dice loss minimizes the discrepancy between the predicted and ground truth masks, ensuring accurate lesion segmentation.

$$\mathcal{L}_{\text{Dice}} = 1 - 2 \times \frac{\sum_p y_p \times \hat{y}_p}{\sum_p y_p + \sum_p \hat{y}_p} \quad (4)$$

One of the key advantages of U-Net is its reduced complexity compared with other segmentation models, such as DeepLab or Mask R-CNN. This reduction in complexity not only simplifies the training process but also reduces latency, making the model more suitable for real-time applications⁵². Furthermore, the lesions in the HAM10000 dataset are generally large enough to be effectively analyzed via a moderately complex model such as U-Net⁵³.

Employing an overly complex and advanced model for this task would be computationally inefficient and unnecessary, as the additional complexity would not yield significant improvements in segmentation accuracy. Instead, U-Net strikes an optimal balance between performance and computational efficiency, making it an ideal choice for this study.

The output of the U-Net model is a binary segmentation mask that precisely delineates the lesion region. This mask then isolates the lesion from the background, enabling focused feature extraction and classification in subsequent pipeline stages. By utilizing U-Net's ability to capture fine-grained details while maintaining computational efficiency, this study ensures accurate and efficient lesion segmentation, laying the foundation for robust skin lesion diagnosis.

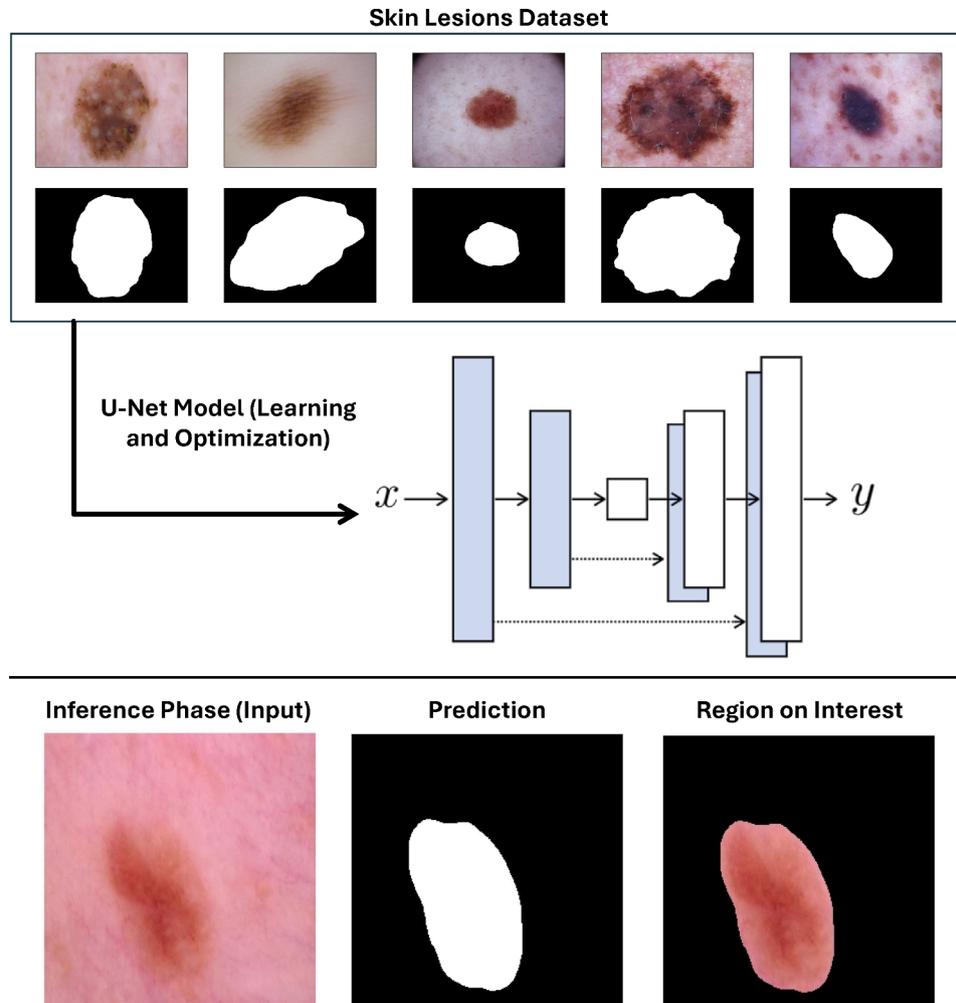


Fig. 2. Schematic of the utilized U-Net model showing the training and inference phases.

Dual-stream feature extraction

Recent advancements in transformer-based architectures, such as Vision Transformers (ViTs) and Swin Transformers^{54–56}, have demonstrated superior performance in capturing long-range dependencies within images, making them particularly well-suited for medical image analysis⁵⁷. In line with this, the proposed dual-stream framework utilizes transformer-based models for both feature extraction streams.

Specifically, Virchow2, a self-supervised Vision Transformer pretrained on 3.1 million whole-slide histopathology images, is employed to extract high-level histopathological features. Its architecture inherently excels at capturing intricate patterns and subtle morphological characteristics of skin lesions, enabling state-of-the-art performance in various computational pathology tasks^{58,59}. Similarly, Nomic, the second stream's feature extractor, utilizes a vision-based transformer architecture to analyze spatial and contextual information from dermatoscopic images⁶⁰.

By integrating these transformer-based models, the framework ensures optimal extraction of both global and local features, addressing the reviewer's recommendation to enhance the capture of subtle lesion characteristics. This design choice underscores the robustness of the proposed approach in overcoming the challenges associated with skin lesion diagnosis.

Virchow2 incorporates a multilayer perceptron (MLP) with gated linear unit (GLU) style gating and sigmoid linear unit (SiLU) activation functions. The GLU mechanism allows for dynamic feature selection, enhancing the model's ability to focus on relevant patterns in the histopathological data. Register tokens are ignored during processing, ensuring that only meaningful patch and class tokens contribute to the final embeddings.

The embeddings are derived from both class tokens c and patch tokens P , which are concatenated to form a comprehensive feature representation as in Equation 5, where $c \in \mathbb{R}^d$ is the class token, $P_i \in \mathbb{R}^d$ represents the i -th patch token, N is the number of patches, and \parallel denotes concatenation. This ensures that the model captures both the global and local features critical for accurate diagnosis.

$$E_{\text{Virchow2}} = \left[c \parallel \frac{1}{N} \times \sum_{i=1}^N P_i \right] \quad (5)$$

The second track employs Nomic, a vision-based feature extraction model, to analyze spatial and contextual information from the same set of images. Nomic processes the images through a transformer-based architecture, extracting embeddings $E_{\text{Nomic}} \in \mathbb{R}^d$ that encode visual patterns such as texture, color, and structural irregularities. The Nomic model uses a Nomic Processor to preprocess the input images, followed by a transformer backbone that generates a last hidden state. The class token from this hidden state is extracted and normalized via L2 normalization, as defined in Equation 6, to ensure consistent scaling of the feature vectors where $h_{\text{cls}} \in \mathbb{R}^d$ represents the class token extracted from the last hidden state of the transformer.

$$E_{\text{Nomic}} = \frac{h_{\text{cls}}}{\|h_{\text{cls}}\|_2} \quad (6)$$

Combining these embeddings with the histopathological-inherited features from Virchow2 provides the model with a more holistic understanding of the lesion characteristics, enhancing its diagnostic capabilities. The fusion process integrates global and local histopathological features with spatial and contextual visual patterns, enabling the model to achieve superior performance in skin lesion classification.

Feature fusion and integration

The features extracted from both tracks are integrated through a fusion mechanism that combines the strengths of histopathologically inherited and vision-based data. The embeddings from Virchow2 and Nomic are normalized and concatenated, ensuring that the model retains the discriminative power of both feature sets. The fusion process can be expressed as in Equation 7, where $\text{Norm}(\cdot)$ denotes L2 normalization.

$$E_{\text{fused}} = \text{Norm}(E_{\text{Virchow2}}) \parallel \text{Norm}(E_{\text{Nomic}}) \quad (7)$$

$$\text{Norm}(x) = \frac{x}{\|x\|_2} \quad (8)$$

This fusion process is followed by a series of fully connected (FC) dense layers, which further refine the combined feature representation. Each dense block consists of an FC layer, a LeakyReLU activation function, and a dropout layer to prevent overfitting. Integrating these features is critical for capturing the complex interplay between morphological and visual characteristics, enabling the model to make more informed decisions during classification.

Classification model

The classification model is built upon a deep neural network architecture incorporating fused feature representations. The model consists of a multilayer perceptron (MLP) with dropout and batch normalization layers to prevent overfitting and improve generalizability. The output logits z are computed via Equation 9, where W_1 , W_2 are weight matrices, b_1 , b_2 are bias terms, and ReLU is the activation function.

$$z = W_2 \times \text{ReLU}(W_1 \times E_{\text{fused}} + b_1) + b_2 \quad (9)$$

The final probabilities p are obtained via the Softmax function presented in Equation 10, where C is the number of classes.

$$p_i = \frac{\exp(z_i)}{\sum_{j=1}^C \exp(z_j)} \quad (10)$$

The model is trained via a cross-entropy loss function, as expressed in Equation 11, where y_i is the ground truth label. The inclusion of both histopathologically inherited and vision-based features allows the model to achieve high classification accuracy while maintaining robustness to variations in the input data.

$$\mathcal{L} = - \sum_{i=1}^C y_i \times \log(p_i) \quad (11)$$

To ensure efficient training and convergence, the learning rate was dynamically adjusted using a ReduceLROnPlateau scheduler. This scheduler reduces the learning rate when the validation loss plateaus, as defined in Equation 12, where η_t is the learning rate at epoch t , η_{t-1} is the learning rate at the previous epoch, and α is the reduction factor applied when the validation loss stops improving.

$$\eta_t = \begin{cases} \eta_{t-1} & \text{if } \mathcal{L}_{\text{val}}(t) < \mathcal{L}_{\text{val}}(t-1), \\ \alpha \times \eta_{t-1} & \text{if } \mathcal{L}_{\text{val}}(t) \geq \mathcal{L}_{\text{val}}(t-1). \end{cases} \quad (12)$$

This approach effectively balances computational efficiency and model performance, ensuring stable convergence without excessive computational overhead. While techniques such as Cosine Annealing or OneCycleLR can

be potential alternatives for improving convergence speed and avoiding local minima, the current strategy already demonstrates robust performance across multiple trials. Nevertheless, future suggestions is to explore the integration of these advanced learning rate schedules to further optimize the model's training dynamics and enhance its adaptability to complex datasets.

Model evaluation and validation

The performance of the proposed model is evaluated via a comprehensive set of metrics, including precision, recall, F1 score, accuracy, and specificity. These metrics are computed via weighted averaging techniques to account for class imbalance in the dataset⁶¹. For example, weighted precision $\text{Precision}_{\text{weighted}}$ is calculated via Equation 13, where $w_i = \frac{n_i}{N}$ is the weight for class i , n_i is the number of samples in class i , and N is the total number of samples.

$$\text{Precision}_{\text{weighted}} = \sum_{i=1}^C w_i \times \frac{\text{TP}_i}{\text{TP}_i + \text{FP}_i} \quad (13)$$

Similarly, the weighted F1 score is computed via Equation 14, where precision_i and recall_i are the precision and recall for class i , respectively⁶².

$$\text{F1}_{\text{weighted}} = \sum_{i=1}^C w_i \times \frac{2 \times \text{Precision}_i \times \text{Recall}_i}{\text{Precision}_i + \text{Recall}_i} \quad (14)$$

Weighted accuracy is defined as in Equation 15, where TN_i and FN_i are the true negatives and false negatives for class i , respectively.

$$\text{Accuracy}_{\text{weighted}} = \sum_{i=1}^C w_i \times \frac{\text{TP}_i + \text{TN}_i}{\text{TP}_i + \text{TN}_i + \text{FP}_i + \text{FN}_i} \quad (15)$$

The model is validated through a series of experiments, including k-fold cross-validation and independent testing on unseen data. Additionally, dimensionality reduction techniques such as PCA, t-SNE, and UMAP are employed to visualize the feature embeddings, providing insights into the model's ability to separate different classes in the latent space. For example, PCA reduces the dimensionality of the embeddings $E_{\text{fused}} \in \mathbb{R}^d$ to $E_{\text{PCA}} \in \mathbb{R}^2$ via Equation 16, where U and Σ are the eigenvectors and eigenvalues of the covariance matrix of E_{fused} .

$$E_{\text{PCA}} = U \times \Sigma \quad (16)$$

The confusion matrix is used to analyze the model's performance at the granular level, identifying potential areas for improvement. The results demonstrate that the dual-stream approach significantly enhances diagnostic accuracy compared to single-track models, making it a promising solution for detecting skin cancer.

To illustrate the feature extraction and fusion process, consider a skin lesion image I . Virchow2 first processes the image to extract histopathologically inherited features E_{Virchow2} , and Nomic is used to extract vision-based features E_{Nomic} . These features are normalized and concatenated to form the fused embedding E_{fused} . The fused embedding is then passed through the classification model to produce the final probabilities p , as shown in the following flow:

1. Input Image: I
2. Feature Extraction:
 - Virchow2: $E_{\text{Virchow2}} = \left[c \parallel \frac{1}{N} \times \sum_{i=1}^N P_i \right]$
 - Nomic: E_{Nomic}
3. Feature Fusion: $E_{\text{fused}} = \text{Norm}(E_{\text{Virchow2}}) \parallel \text{Norm}(E_{\text{Nomic}})$
4. Classification: $z = W_2 \times \text{ReLU}(W_1 \times E_{\text{fused}} + b_1) + b_2$
5. Output Probabilities: $p_i = \frac{\exp(z_i)}{\sum_{j=1}^C \exp(z_j)}$

The proposed framework pseudocode

The pseudocode for the proposed framework is outlined in Algorithm 1. The algorithm provides a step-by-step description of the dual-stream deep learning framework for skin lesion diagnosis, encompassing data preprocessing, feature extraction using Virchow2 and Nomic, feature fusion, model training, and performance evaluation. Each step is designed to ensure robustness, generalizability, and interpretability of the model while addressing challenges such as class imbalance and complex lesion characteristics. The modular structure of the pseudocode reflects the key components of the framework, enabling reproducibility and facilitating future enhancements.

```

Input: HAM10000 dataset  $D$ , Pre-trained models: Virchow2, Nomic, U-Net
Output: Trained classification model  $M^*$ , Performance metrics, Visualizations
// Step 1: Data Preprocessing
1 foreach image  $I \in D$  do
  // Resize image to  $224 \times 224$  pixels.
2    $I_{\text{resized}} \leftarrow \text{Resize}(I)$ 
  // Apply data augmentation techniques (rotation, flipping, scaling, color jittering).
3    $I_{\text{augmented}} \leftarrow \text{Augment}(I_{\text{resized}})$ 
  // Normalize pixel intensities to  $[0, 1]$ .
4    $I_{\text{normalized}} \leftarrow \text{Normalize}(I_{\text{augmented}})$ 
  // Perform lesion segmentation using U-Net.
5    $\text{ROI} \leftarrow \text{U-NetSegmentation}(I_{\text{normalized}})$ 
  // Store preprocessed images and ROIs.
6    $D_{\text{preprocessed}}[I] \leftarrow (\text{ROI}, I_{\text{normalized}})$ 
// Step 2: Feature Extraction
7 foreach preprocessed image  $(\text{ROI}, I_{\text{normalized}}) \in D_{\text{preprocessed}}$  do
  // Extract histopathologically inherited features using Virchow2.
8    $E_{\text{Virchow2}} \leftarrow \text{ExtractEmbedding}(\text{ROI}, \text{Virchow2})$ 
  // Extract vision-based features using Nomic.
9    $E_{\text{Nomic}} \leftarrow \text{ExtractEmbedding}(I_{\text{normalized}}, \text{Nomic})$ 
  // Concatenate embeddings from both streams.
10   $E_{\text{fused}} \leftarrow \text{Concat}(E_{\text{Virchow2}}, E_{\text{Nomic}})$ 
  // Store fused embeddings for classification.
11   $Z[I] \leftarrow E_{\text{fused}}$ 
// Step 3: Model Training and Optimization
12 foreach classifier  $C \in \{\text{MLP}, \text{CNN}, \text{Ensemble}\}$  do
  // Split the fused embeddings into training, validation, and test sets.
13   $Z_{\text{train}}, Z_{\text{val}}, Z_{\text{test}} \leftarrow \text{Split}(Z)$ 
  // Perform hyperparameter tuning for the classifier.
14   $\theta^* \leftarrow \text{HyperparameterTuning}(C, Z_{\text{train}}, Z_{\text{val}})$ 
  // Train the classifier with optimal hyperparameters.
15   $M \leftarrow \text{Train}(C, Z_{\text{train}}, \theta^*)$ 
  // Evaluate the classifier on the validation set.
16   $\text{Performance} \leftarrow \text{Evaluate}(M, Z_{\text{val}})$ 
  // Store the best-performing model.
17  if  $\text{Performance} > \text{BestPerformance}$  then
18     $M^* \leftarrow M$ 
19     $\text{BestPerformance} \leftarrow \text{Performance}$ 
// Step 4: Performance Evaluation
// Evaluate the best-performing model on the test set.
20  $\text{Metrics} \leftarrow \text{Evaluate}(M^*, Z_{\text{test}})$ 
  // Calculate confidence intervals for performance metrics.
21  $\text{ConfidenceIntervals} \leftarrow \text{CalculateConfidenceIntervals}(\text{Metrics})$ 
  // Perform statistical significance testing.
22  $\text{Significance} \leftarrow \text{StatisticalTest}(M^*, Z_{\text{test}})$ 
// Step 5: Generate Visualizations for Interpretation and Analysis
23  $\text{PCAPlot} \leftarrow \text{PlotPCA}(Z)$ 
24  $\text{ConfusionMatrix} \leftarrow \text{PlotConfusionMatrix}(M^*, Z_{\text{test}})$ 
25  $\text{ROCCurve} \leftarrow \text{PlotROCCurve}(M^*, Z_{\text{test}})$ 
26  $\text{PrecisionRecallCurve} \leftarrow \text{PlotPrecisionRecallCurve}(M^*, Z_{\text{test}})$ 

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Algorithm 1. The proposed dual-stream framework for skin lesion diagnosis.

Experiments and discussion

The HAM10K dataset, which consists of 10,015 dermatoscopic images across seven common skin lesion types, was utilized in this study. To evaluate the proposed dual-stream framework, 70% of the dataset was used for training, 15% for validation, and the remaining 15% for testing. This split ensures a robust assessment of the model's generalization capabilities on unseen data. The performance was assessed via a comprehensive set of metrics, including precision, recall, F1 score, accuracy, and specificity.

Segmentation performance

The U-Net architecture was employed for precise lesion segmentation, ensuring accurate localization of affected areas. Figure 3 shows the learning and validation curves for segmentation, including metrics such as loss, accuracy, Dice coefficient, intersection over union (IoU), recall, precision, and other key performance indicators. The curves indicate stable convergence during training, with minimal overfitting, demonstrating the model's ability to generalize well to unseen data.

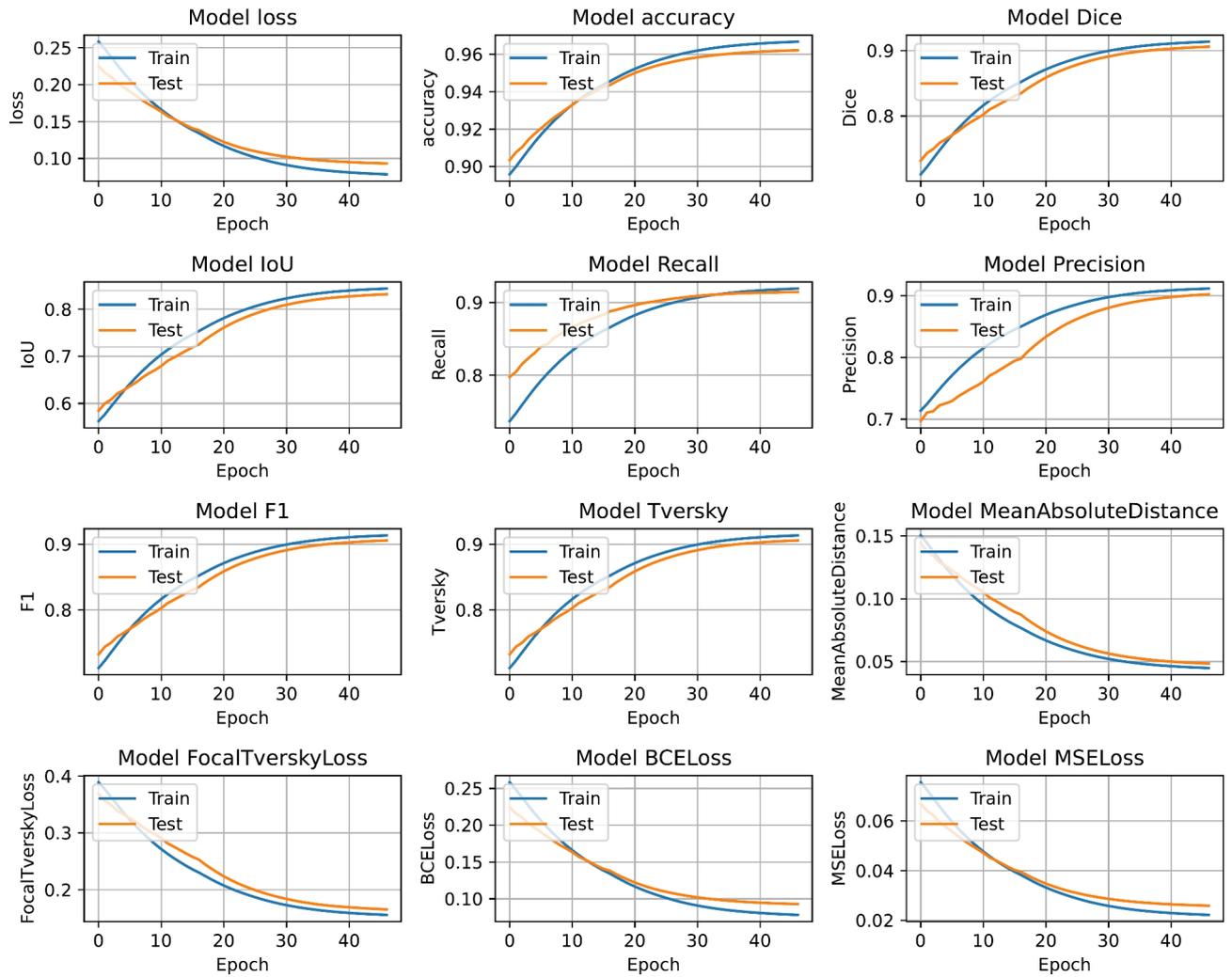


Fig. 3. Training and validation curves for the U-Net segmentation model. The plots depict the loss, accuracy, Dice coefficient, intersection over union (IoU), recall, and precision over epochs, demonstrating stable convergence and robust performance.

Metric	Value
BCE Loss	0.0754
Dice Coefficient	0.9169
F1 Score	0.9169
Focal Tversky Loss	0.1514
IoU	0.8493
MSE Loss	0.0213
Mean Absolute Distance	0.0432
Precision	0.9146
Recall	0.9225
Tversky Index	0.9169
Accuracy	96.79%
Overall Loss	0.0754

Table 1. Quantitative results for U-Net segmentation.

Quantitative results from the segmentation process are summarized in Table 1, showcasing the model's robust performance across multiple evaluation metrics. Notably, the model achieved a Dice coefficient of 0.9169, an IoU of 0.8493, and a mean accuracy of 96.79%, reflecting its effectiveness in delineating lesion boundaries even in challenging cases with irregular shapes and textures. Additionally, the precision and recall values of 0.9146 and 0.9225, respectively, highlight the model's balanced performance in identifying both positive and negative regions. The Focal Tversky Loss and BCELoss were minimized to 0.1514 and 0.0754, respectively, further confirming the model's strong optimization during training.

Figure 4 shows three prediction samples from the U-Net model. The columns represent the original dermatoscopic images, ground truth masks, and predicted masks, respectively. The visual results highlight the model's ability to accurately segment skin lesions, even in cases with complex boundaries and varying textures. For example, the model successfully identifies lesions with heterogeneous pigmentation and irregular edges, which are often challenging for traditional segmentation methods. This level of precision is crucial for downstream tasks such as feature extraction and classification, as accurate segmentation ensures that the extracted features represent the lesion's true characteristics.

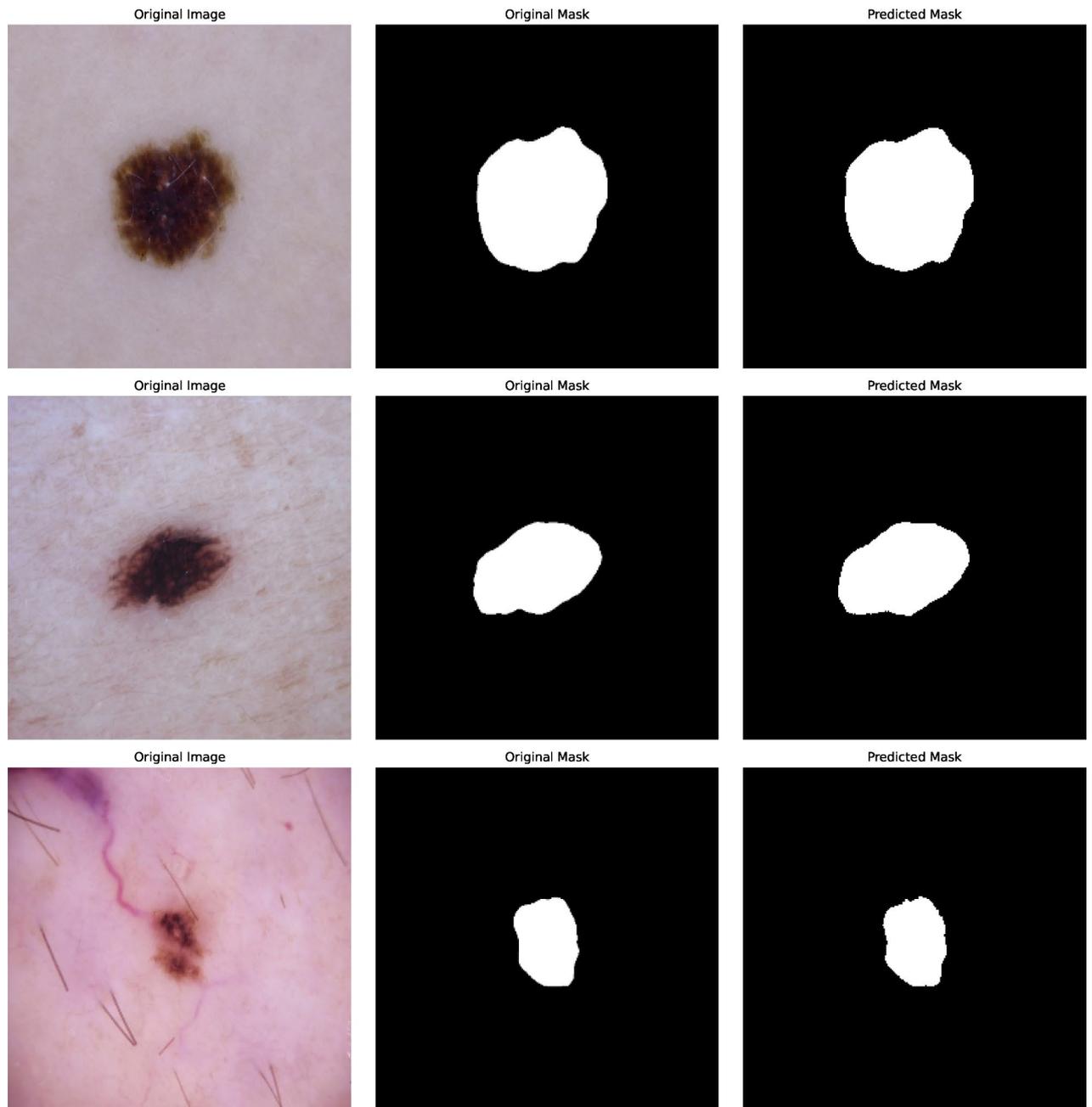


Fig. 4. Visualization of U-Net segmentation predictions. The columns display (from left to right) the following: original dermatoscopic images, ground truth masks, and predicted masks. The results highlight the model's ability to segment skin lesions with complex boundaries and varying textures accurately.

Trials	Precision	Recall	F1	Accuracy	Specificity
Trial 1	93.16%	93.21%	93.19%	95.85%	91.37%
Trial 2	93.80%	93.88%	93.84%	96.29%	92.08%
Trial 3	94.20%	94.21%	94.20%	96.39%	91.87%
Trial 4	93.43%	93.48%	93.45%	96.30%	92.33%
Trial 5	94.20%	94.21%	94.21%	96.60%	92.66%
Trial 6	93.97%	93.88%	93.93%	96.16%	91.10%
Trial 7	94.12%	94.15%	94.13%	96.57%	92.37%
Trial 8	94.27%	94.21%	94.24%	96.55%	92.54%
Trial 9	93.62%	93.61%	93.62%	95.73%	90.52%
Trial 10	93.07%	93.15%	93.11%	96.10%	91.39%
Max	94.27%	94.21%	94.24%	96.60%	92.66%
Mean	93.78%	93.80%	93.79%	96.25%	91.82%
Std	0.00444	0.00415	0.00428	0.00298	0.00704
CI	0.00275	0.00257	0.00266	0.00184	0.00437

Table 2. Performance metrics of the proposed dual-stream approach on the HAM10K dataset. The results are reported over 10 trials, including precision, recall, F1 score, accuracy, and specificity. The maximum, mean, standard deviation (Std), and confidence interval (CI) values are also provided.

Study	Precision	Recall	F1	Accuracy	Specificity
Removal of Embedding	61.33%	66.93%	64.01%	75.02%	33.07%
Removal of Vision	85.51%	85.48%	85.50%	88.43%	82.81%

Table 3. Performance metrics of ablation studies evaluating the contribution of each component in the dual-stream framework. The results are reported for scenarios where either the embedding branch (Virchow2) or the vision branch (Nomic) is removed.

Classification performance

The classification performance of the proposed dual-stream approach is summarized in Table 2. The results of over 10 trials were reported, with metrics including precision, recall, F1 score, accuracy, and specificity. The maximum, mean, standard deviation (Std), and confidence interval (CI) values are also provided. The model achieves a mean accuracy of 96.25% and a mean F1 score of 93.79%, demonstrating its robustness and consistency across trials. The high specificity values (mean of 91.82%) further indicate the model's ability to correctly identify negative cases, which is critical for reducing false positives in skin cancer diagnosis.

The precision and recall values, which are particularly important for medical applications, are consistently high across all trials. This indicates that the model strikes a balance between minimizing false positives (high precision) and false negatives (high recall), ensuring reliable diagnostic outcomes. The low standard deviation and narrow confidence intervals for all the metrics further underscore the stability and reproducibility of the proposed approach.

Ablation studies

Ablation studies were conducted to evaluate the contribution of each component in the dual-stream framework. Table 3 presents the performance metrics when either the embedding branch (Virchow2) or the vision branch (Nomic) is removed. The results show a significant decrease in performance when the embedding branch is removed, with the precision and F1 score decreasing to 61.33% and 64.01%, respectively. This highlights the importance of histopathologically inherited features in capturing domain-specific patterns for accurate diagnosis. The embedding branch, which uses Virchow2, provides high-level morphological information critical for distinguishing between different skin lesions.

Similarly, removing the vision branch results in a noticeable decline in performance, with precision and the F1 score decreasing to 85.51% and 85.50%, respectively. This underscores the complementary role of vision-based features in enhancing the model's diagnostic capabilities. The vision branch, powered by Nomic, captures spatial and contextual information such as texture, color, and structural irregularities, which are essential for identifying subtle differences between lesion types. The ablation studies demonstrate that both branches contribute uniquely to the model's performance, and their integration is key to achieving state-of-the-art results.

Figure 5 visualizes the dimensionality reduction of the Virchow2 features via principal component analysis (PCA). The plot reflects the complexity of the problem and provides insights into the separability of different lesion classes in the feature space. The PCA visualization shows distinct clusters for different lesion types, indicating that the Virchow2 embeddings effectively capture discriminative features. However, some overlap between classes is observed, highlighting the challenges of distinguishing between visually similar lesions. This

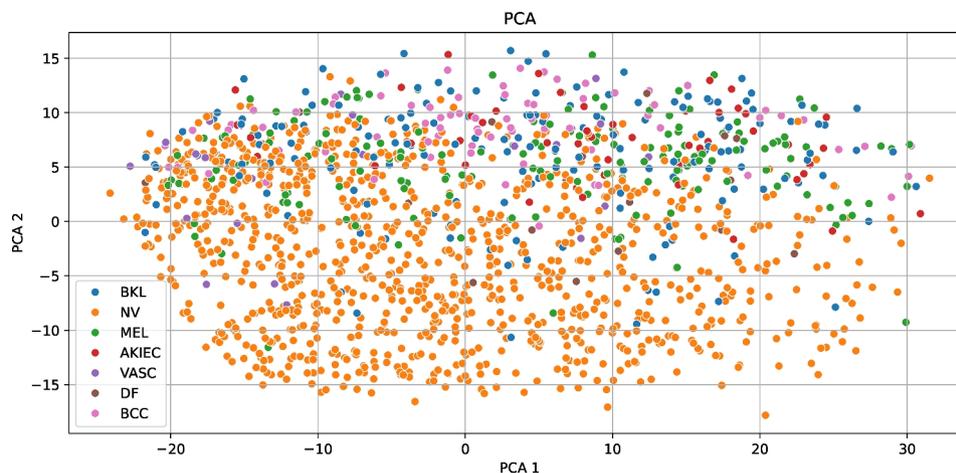


Fig. 5. Dimensionality reduction of Virchow2 embeddings via principal component analysis (PCA). The plot illustrates the separability of different lesion classes in the feature space, highlighting the complexity of the problem and the importance of combining histopathologically inherited and vision-based features.

Study (Ref)	Year	Approach	Results (Accuracy)
Chaturvedi et al. ⁶³	2021	MobileNet	An overall accuracy of 83.1% for seven classes
Xin et al. ⁴⁴	2022	Improved transformer network	94.3% accuracy
Agyenta et al. ⁶⁴	2022	DenseNet201	Accuracy of 99.12% for training and 86.91% for testing
Ali et al. ⁴¹	2022	EfficientNets	An F1 score of 87% and a top-1 accuracy of 87.91%
Adebiyi et al. ⁴³	2024	Multimodal learning	Best accuracy (94.11%) and AUCROC (0.9426)
Proposed Dual-Stream	2025	U-Net + Virchow2 + Nomic	96.25% (See Table 2)

Table 4. Comparison of the proposed dual-stream approach with existing methods using the HAM10000 dataset.

further emphasizes the importance of combining histopathologically inherited and vision-based features, as the dual-stream approach utilizes complementary information to improve class separability.

Comparative analysis

The proposed dual-track approach outperforms traditional single-track methods, which rely solely on either extracted or vision-based features. Integrating both feature types enables the model to capture a more comprehensive representation of skin lesions, improving diagnostic accuracy. For example, while extracted features effectively identify morphological patterns, they may struggle with lesions that exhibit similar structures but differ in texture or color. Conversely, vision-based features excel at capturing visual patterns but may lack the domain-specific knowledge required for precise diagnosis. By combining these two modalities, the proposed approach addresses the limitations of single-track methods and achieves superior performance.

To contextualize the performance of the proposed dual-stream approach, a comparative analysis with existing studies using the HAM10000 dataset is presented in Table 4. It highlights the methodologies, results, and key metrics of recent studies alongside the performance of the proposed framework.

The proposed dual-stream approach achieves an accuracy of 96.25%, outperforming several recent studies. For example, Chaturvedi et al.⁶³ employed MobileNet for skin lesion classification, achieving an accuracy of 83.1%. While their approach is computationally efficient, it lacks the ability to integrate domain-specific features, which limits its ability to distinguish between visually similar lesions. Similarly, Xin et al.⁴⁴ proposed an improved transformer network, achieving an accuracy of 94.3%. Although their method utilizes advanced transformer architectures, it does not explicitly incorporate embeddings, which are critical for capturing morphological patterns.

Agyenta et al.⁶⁴ utilized DenseNet201, reporting a training accuracy of 99.12% but a testing accuracy of only 86.91%. This significant drop in performance suggests potential overfitting, highlighting the challenges of generalizing single-modality approaches to unseen data. In contrast, the proposed dual-stream approach demonstrates robust generalization, as evidenced by its consistent performance across multiple trials (see Table 2).

Ali et al.⁴¹ explored EfficientNets for multiclass skin lesion classification, achieving an F1 score of 87% and a top-1 accuracy of 87.91%. While their approach is effective, it relies solely on vision-based features, which may not fully capture the complexity of skin lesions. The proposed dual-stream framework addresses this limitation by integrating histopathologically inherited features, resulting in higher accuracy and F1 scores.

Adebiyi et al.⁴³ introduced a multimodal learning approach, achieving an accuracy of 94.11% and an AUCROC of 0.9426. Their method combines multiple data modalities, demonstrating the benefits of integrating diverse feature types. However, their approach does not explicitly utilize pretrained models such as Virchow2, which are specifically designed for dermatoscopic data. The proposed dual-stream approach builds on this idea by incorporating Virchow2 embeddings, further enhancing diagnostic accuracy.

Clinical implications

The high accuracy and robustness of the proposed framework have significant implications for clinical practice. Skin cancer diagnosis often requires expert dermatologists to analyze complex visual and morphological patterns, which can be time-consuming and prone to human error. The proposed dual-stream approach provides a reliable and efficient tool for assisting clinicians in diagnosing skin lesions, reducing the burden on healthcare systems and improving patient outcomes. The interpretability of the model, as demonstrated by PCA visualization and ablation studies, further enhances its clinical utility, as it allows clinicians to understand the basis for the model's predictions.

Moreover, the model's ability to generalize across different trials and datasets suggests that it can be adapted to various clinical settings and populations. This is particularly important for skin cancer diagnosis, as lesion characteristics can vary significantly depending on factors such as skin type, age, and geographic location. The robustness of the proposed framework to such variations makes it a promising candidate for widespread deployment in dermatology.

Limitations

Despite its strong performance, the proposed approach has several limitations. First, the model relies on high-quality dermatoscopic images, which may not always be available in resource-limited settings. Future work could explore alternative imaging modalities or techniques for enhancing low-quality images. Second, while ablation studies demonstrate the importance of both feature types, further research is needed to optimize the fusion mechanism and explore alternative architectures for integrating histopathologically inherited and vision-based features.

Additionally, the model's performance could be further improved by incorporating additional data sources, such as patient metadata (e.g., age, sex, medical history) or multispectral imaging data. These extra inputs could provide valuable context for improving diagnostic accuracy and personalizing treatment plans. Finally, future work should validate the model on more extensive and diverse datasets to ensure its generalizability across different populations and clinical scenarios.

Conclusions and future directions

This study presents a dual-stream DL framework for skin lesion diagnosis that combines histopathological-inherited and vision-based feature extraction to achieve state-of-the-art performance. The proposed approach uses advanced DL techniques to address key limitations of traditional diagnostic methods, such as subjectivity, interobserver variability, and resource constraints. The U-Net architecture ensures precise lesion segmentation, whereas the dual-stream feature extraction mechanism, which uses Virchow2 and Nomic, captures both the morphological and contextual information critical for accurate diagnosis. The fusion of these features enables the model to achieve a mean accuracy of 96.25% and a mean F1 score of 93.79% on the HAM10000 dataset, demonstrating its robustness and generalizability across multiple trials. Ablation studies highlight the complementary roles of histopathological and vision-based streams, with the removal of either stream leading to significant performance degradation. Comparative analysis with existing studies further validates the superiority of the proposed framework, which outperforms traditional single-modality approaches and recent multimodal methods. The interpretability of the model, as demonstrated by dimensionality reduction techniques such as PCA, provides valuable insights into the feature space, enhancing its clinical utility. The proposed framework has significant implications for clinical practice, offering a reliable and efficient tool for assisting dermatologists in diagnosing skin lesions. Its ability to generalize across diverse datasets and populations makes it a promising candidate for widespread deployment, particularly in resource-limited settings.

Future directions

Future work will focus on key enhancements to strengthen the framework. First, the fusion mechanism will be optimized, potentially incorporating attention mechanisms or multi-scale feature fusion to better capture local and global spatial information. Second, additional data sources, such as patient metadata (e.g., age, sex, medical history) and advanced imaging modalities like thermal or multispectral data, multispectral or 3D imaging, will be integrated to enrich contextual information and enable personalized predictions. Third, the model will be validated on larger, diverse datasets, expanding the evaluation to include datasets that explicitly represent underrepresented populations, varied skin phototypes (e.g., Fitzpatrick skin types I-VI), and regional differences in lesion presentation. These improvements aim to revolutionize skin cancer diagnosis, enhance accuracy, improve outcomes, and reduce mortality, paving the way for clinical adoption.

Data Availability

This study used the HAM10000 dataset, a large collection of multi-source dermatoscopic images of common pigmented skin lesions, which is available at: <https://dataverse.harvard.edu/dataset.xhtml?persistentId=doi:10.7910/DVN/DBW86T>

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Declarations

Competing interests

The author declares that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

Ethics approval and consent to participate

Not applicable

Additional information

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