## Highlights

MedDef: An Efficient Self-Attention Model for Adversarial Resilience in Medical Imaging with Unstructured Pruning

E.K. Dongbo, S. Niu, P. Fero, P. Bargin, J.N. Kofa

- Novel Defense-Aware Attention Mechanism (DAAM) integrates adversarial robustness into feature extraction
- Medical domain-aware defensive strategy preserves diagnostic features while suppressing attacks
- Unstructured pruning enhances security rather than compromising it in medical imaging
- $\bullet$  Achieves 97.52% adversarial accuracy with maintained diagnostic performance
- Comprehensive evaluation on Retinal OCT and Chest X-Ray datasets against multiple attacks

# MedDef: An Efficient Self-Attention Model for Adversarial Resilience in Medical Imaging with Unstructured Pruning

E.K. Dongbo<sup>a</sup>, S. Niu<sup>a,\*</sup>, P. Fero<sup>b</sup>, P. Bargin<sup>b</sup>, J.N. Kofa<sup>c</sup>

<sup>a</sup>School of Information Science and Engineering, University of Jinan, , Jinan, 250022, Shandong, P.R. China <sup>b</sup>School of Computer Science & Technology, Zhejiang Sci-Tech University, , Hangzhou, 310018, , P.R. China <sup>c</sup>College of Informatics, Huazhong Agricultural University, , Wuhan, 430070, , P.R. China

#### Abstract

In an effort to improve diagnostic precision, medical imaging systems are increasingly incorporating artificial intelligence (AI). However, these systems remain susceptible to adversarial attacks, which are subtle, undetectable disruptions intended to trick models into generating inaccurate results. While current methods like adversarial training and input preprocessing provide some partial answers, they frequently reduce diagnostic accuracy by not differentiating between adversarial noise and fine-grained signals that are medically important. We introduce Medical Defense (MedDef), a novel defensive architecture that addresses this challenge by integrating a Defense-Aware Attention Mechanism (DAAM) with unstructured pruning to achieve robust adversarial resilience. DAAM signifies a transition from post-hoc defenses to an integrated approach to robustness, comprising three interrelated components: Adversarial Feature Detection (for noise suppression), Medical Feature Extraction (for domain-specific feature enhancement), and Multi-Scale Feature Analysis (for coordinated multi-resolution defense). These components collaboratively identify and neutralize adversarial noise while amplifying diagnostically critical features. Extensive experiments on Retinal OCT and Chest X-Ray datasets against four common attack methods demonstrate that MedDef achieves exceptional robustness (up to 97.52% adversarial accuracy)

Email address: sjniu@hotmail.com (J.N. Kofa)

<sup>\*</sup>Corresponding author

while maintaining high diagnostic accuracy, establishing that security and diagnostic performance can be simultaneously optimized rather than traded off, laying the foundation for clinically viable, adversarially robust medical imaging systems.

Keywords: Adversarial Resilience, Medical Imaging, Defense-Aware Attention Mechanism, Unstructured Pruning, Robust Model

#### 1. Introduction

Deep neural networks have revolutionized medical imaging analysis, achieving unprecedented diagnostic accuracy across various conditions [1]. While these systems approach or exceed human-level performance in specialized tasks, they remain vulnerable to adversarial attacks; imperceptible perturbations that cause incorrect predictions with potentially serious clinical consequences [2, 3].

Current defense strategies fall into several areas, which can be categorically grouped into three: (1) input preprocessing techniques (denoising [4], JPEG compression [5]) that neutralize perturbations; (2) model regularization approaches like adversarial training [6] that improve robustness; and (3) architectural modifications (defensive distillation [7], feature squeezing [8], ensemble methods [9]) that detect or mitigate adversarial inputs. The success of these methods is, however, constrained by the particular difficulties associated with medical imaging. Research shows adversarial training reduced classification accuracy for subtle pathological features by 8-12% [10], while preprocessing techniques decreased sensitivity to critical diagnostic signals by up to 14% [11]. Comparative studies across modalities revealed state-of-theart defenses reduce clear image performance by 5-16% [12], highlighting an unacceptable robustness-accuracy tradeoff.

Three critical challenges emerge in medical imaging defense that directly motivate our DAAM design:

Challenge 1: Noise Suppression vs. Feature Preservation. Preprocessing procedures typically suppress delicate, localized textures and patterns that constitute diagnostic features in medical images [4]. Early retinal pathology, for example, shows up as microscopic alterations in the thickness of the photoreceptor layer, whereas antagonistic perturbations take advantage of comparable high-frequency features. Our Adversarial Feature Detection (AFD) component targets noise patterns while maintaining diagnostic sub-

tleties because a defense mechanism that can distinguish between adversarial noise and medically relevant fine-grained characteristics is required.

Challenge 2: Vulnerability Dependent on Scale. From cellular-level anomalies to organ-level structural alterations, medical pictures provide diagnostically important information at several spatial dimensions. By focusing on particular resolution levels where defenses are weakest, adversarial attacks take advantage of this multi-scale nature [13]. Research in dermatology showed that feature squeezing improved robustness by 23% but decreased the sensitivity of early melanoma detection by 17%, emphasizing the necessity of scale-aware protection. Our Multi-Scale Feature Analysis (MSF) component is directly motivated by this to offer coordinated defense across all pertinent spatial resolutions.

Challenge 3: Robustness Requirements Specific to Domains. Anatomically significant characteristics (edges, textures, morphological patterns) that differentiate healthy tissue from diseased tissue must be preserved for medical imaging. Instead of using medical domain knowledge, conventional defenses treat robustness as a rival to accuracy [10]. This drives our Medical Feature Extraction (MFE) component, which suppresses adversarial perturbations while explicitly enhancing diagnostically significant anatomical characteristics.

To tackle adversarial vulnerabilities in medical imaging, we propose MedDef, a framework built around a Defense-Aware Attention Mechanism (DAAM) that integrates defense directly into feature processing. DAAM comprises three components: Adversarial Feature Detection (AFD), which suppresses high-frequency adversarial noise while preserving fine-grained diagnostic features; Medical Feature Extraction (MFE), which enhances edge and texture cues using domain knowledge; and Multi-Scale Feature Analysis (MSF), which defends across spatial resolutions to maintain robust hierarchical representations. These modules are unified through self-attention that dynamically adjusts feature relevance, enabling DAAM to focus on diagnostically important regions while mitigating attacks. Combined with unstructured pruning, MedDef yields a compact, resilient architecture.

Our contributions include:

1. **Principled DAAM Architecture:** A novel Defense-Aware Attention Mechanism that integrates adversarial robustness directly into feature extraction through three synergistic components (AFD, MFE, MSF), each specifically designed to address distinct vulnerabilities in medical

- adversarial defense. In contrast to post-hoc defenses, DAAM essentially changes the way features are processed in order to optimize both adversary robustness and diagnostic accuracy at the same time.
- 2. Medical Domain-Aware Defensive Strategy: In order to guarantee that defensive capabilities improve rather than degrade diagnostic performance, this defensive strategy is the first to specifically integrate medical imaging domain knowledge (edge detection, texture analysis, and multi-scale pathology) into adversarial defense mechanisms. Our MFE component demonstrates that domain-specific feature enhancement can serve as an effective defensive strategy.
- 3. Compression-Security Synergy: Strategic unstructured pruning was found to be an efficient defensive tactic that counters the widely held belief that robustness is weakened by compression. Our thorough investigation of pruning rates ranging from 0 to 80% indicates optimal compression points that vary by dataset: Chest X-Ray models achieve peak performance at 40% pruning (98.38% vs 97.67% unpruned), while ROCT models maintain optimal performance up to 30% pruning without degradation. This finding establishes that the compression-security relationship is not universally antagonistic but can be leveraged as a defensive mechanism when properly calibrated for medical imaging domains.

#### 2. Related Work

## 2.1. Adversarial Defense Techniques in Medical Imaging

Recent research has increasingly focused on addressing adversarial attacks in medical imaging, which can lead to severe consequences such as misdiagnosis and inaccurate clinical decisions [14]. To solve these issues, a variety of defense tactics have been proposed. These include input pre-processing, adversarial training, and strategies for algorithm comprehension [15]. Zhao [16] presented a strong architecture that enhances resilience against such attacks by combining Unsupervised Adversarial Detection and Semi-Supervised Adversarial Training. Paschali highlighted the importance of evaluating both generalizability and model resilience, demonstrating notable differences in performance in extreme environments [17]. Additionally, Luo proposed a game-theoretic framework integrating conformal prediction to enhance model robustness against both known and unknown adversarial perturbations [18].

Moreover, Alzubaidi introduced the Model Ensemble Feature Fusion (MEFF) technique, which integrates features from many deep learning models to improve robustness against various adversarial attacks across diverse medical imaging applications [9]. Sahu explored the vulnerabilities of deep learning models in medical image diagnosis and proposed adversarial training as a key defense mechanism [13]. These studies collectively highlight the growing importance of developing robust defense strategies to ensure the reliability and accuracy of AI systems in medical imaging. The ongoing developments in this field highlight how critical the need is for continuous innovation to defend medical imaging technologies against adversarial threats. This body of work not only advances our understanding of adversarial resilience but also paves the way for more secure and reliable medical imaging applications in the future [19].

Furthermore, novel techniques have been developed lately to further improve the robustness of medical imaging models. For example, Biswas created a hybrid adversarial training method that enhances model resilience against sophisticated attacks by combining supervised and unsupervised learning techniques [20], and Singh made a significant contribution by using generative adversarial networks (GANs) to generate synthetic adversarial examples that are then used to train and fortify medical imaging models. These innovative methods demonstrate how research in this area is dynamic and always changing, emphasizing the value of interdisciplinary cooperation and continuous development to protect the integrity of medical imaging systems.

Moreover, Dong suggests a brand-new adversarial training strategy that motivates the model to generate output probabilities for an adversarial example that are comparable to those of its "inverse adversarial" counterpart. This was accomplished by carrying out in-depth tests on a range of vision datasets and architectures, which showed that the training approach achieves both natural accuracy among robust models and state-of-the-art robustness. Additionally, it enhances the efficiency of single-step adversarial training methods at a minimal computational cost by employing a universal version of inverse adversarial instances [21].

The vast amount of research in the field of adversarial attacks and defending against these attacks on deep neural networks for medical imaging has yielded a variety of approaches and insightful knowledge. Even though these studies greatly advance our knowledge of medical image analysis, further research is still necessary [22]. In the future, research efforts should con-

centrate on creating methods that tackle the dynamic terrain of adversarial attacks, taking into account fresh situations and possible weaknesses. Furthermore, increasing interpretability and openness and incorporating real-world applications can improve the usefulness and efficiency of defense mechanisms. Sustaining the resilience of deep neural networks and staying ahead of adversarial threats will need continued research as the field develops [23].

#### 2.2. Attack Methods

Adversarial attacks aim to fool machine learning algorithms by subtly altering input images while maintaining their visual integrity. Medical imaging is particularly vulnerable to such attacks, as even small misclassifications can have serious clinical repercussions. Building on these research needs, we evaluate our model using four established attack techniques that represent significant threats to medical imaging systems:

Fast Gradient Sign Method (FGSM) is a single-step, gradient-based approach that perturbs the input by taking the sign of the loss function's gradient [24]. Despite its simplicity, FGSM effectively exposes model vulnerabilities by highlighting the direction in which the loss increases most rapidly.

**Projected Gradient Descent (PGD)** extends FGSM through an iterative process that projects adversarial samples back onto a specified constraint space before adding small perturbations in each iteration [25]. PGD is considered one of the most effective first-order attacks due to its iterative nature, providing a demanding test of model resilience.

Basic Iterative Method (BIM) applies FGSM iteratively with a fixed step size, gradually altering the input over multiple rounds. This approach can progressively degrade model performance, revealing vulnerabilities not apparent with single-step attacks [26].

Jacobian-based Saliency Map Attack (JSMA) utilizes a saliency-based approach that leverages the model's Jacobian to identify the most influential features in the input space. By selectively perturbing these crucial aspects, JSMA creates targeted adversarial examples that particularly challenge defense mechanisms [27].

These techniques, which include both single-step and iterative approaches, provide comprehensive benchmarks for assessing model robustness while maintaining the visual integrity of medical images. Our methodology combines these attack vectors to thoroughly test its resilience against perturbations that could otherwise lead to critical misdiagnoses in clinical applications [28, 29].

## 3. Methodology

This section outline our approach for developing a novel defense-oriented model that synergistically integrates self-attention mechanisms, unstructured pruning, and adversarial training to enhance robustness against adversarial attacks while maintaining high diagnostic accuracy in medical imaging applications.

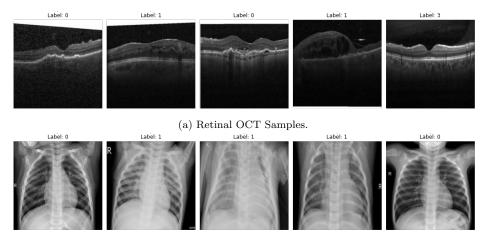
## 3.1. Dataset and Preprocessing

#### 3.1.1. Dataset

Two medical imaging datasets were used: the Retinal OCT (ROCT) dataset (84,484 images across four classes: CNV, DME, Drusen, and Normal) and the Chest X-Ray dataset (5,856 images for binary NORMAL/PNEUMONIA classification). The ROCT dataset was divided into 83,484 training, 32 validation, and 968 test images (242 per class), with original 512×496 grayscale images presenting low brightness and variable aspect ratio challenges. The Chest X-Ray dataset consisted of 4,099 training images (1,108 NORMAL, 2,991 PNEUMONIA), 878 validation images (237 NORMAL, 641 PNEUMONIA), and 879 test images (238 NORMAL, 641 PNEUMONIA), with significant dimension variability (976×544 to 2090×1858) and inconsistent color channels. Both datasets showed variability in image quality, brightness, and contrast that required addressing.

#### 3.1.2. Preprocessing

The preprocessing pipeline standardized both datasets by resizing images to 224×224 pixels with aspect-preserving padding and converting grayscale to three-channel format. For ROCT, we applied min-max scaling to [0,1] followed by standardization with mean [0.19338988] and standard deviation [0.1933612] across channels [30], then enhanced contrast using CLAHE and adjusted brightness. The Chest X-Ray dataset underwent similar resizing and channel conversion, with normalization using dataset-specific mean [0.48230693] and standard deviation [0.22157896] values. We implemented denoising to improve image clarity and removed duplicates (24 identified in the Chest X-Ray dataset) to prevent data leakage [31]. These steps ensured consistently formatted, noise-free images with preserved diagnostic features essential for accurate classification [32].



(b) Chest X-Ray Samples.

Figure 1: Representative medical imaging samples from both datasets: (a) ROCT images showing retinal cross-sections with varying pathological conditions including CNV, DME, Drusen, and Normal cases, and (b) Chest X-Ray images demonstrating normal lung parenchyma and pneumonia infiltrates with diverse imaging characteristics and quality variations.

## 3.2. Proposed Model Architecture and Training Method

#### 3.2.1. Model Architecture

In the development phase of our model, we integrated self-attention mechanisms, unstructured pruning, and adversarial training to enhance both the robustness and efficiency of medical image analysis. The architecture flow advances through three stages; beginning with 1) preprocessing input images from established datasets and applying various data augmentation techniques to diversify the training set; 2) consist of our Defense Aware Attention Mechanism implementation into our model which is then used through stage 3); where we implemented adversarial training approach involves distributing attack perturbations across different data splits: 35% of the training data, 70% of the validation data, and 100% of the test data are exposed to adversarial perturbations. This graded approach to attack exposure ensures thorough model evaluation while maintaining stable training dynamics. Finally, unstructured pruning was done and giving an output of our robust model. The complete architecture is illustrated in Fig. 2.

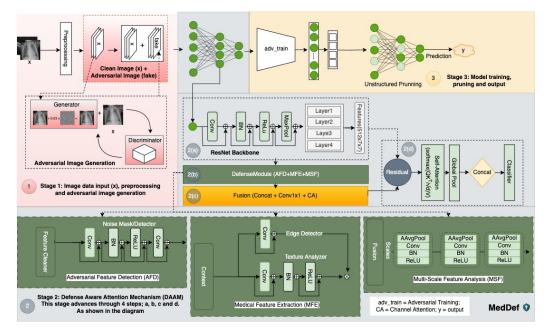


Figure 2: Illustrate the MedDef framework, advancing through 3 stages with stage 1) consisting of the input, processing and adversarial image generation; stage 2) consisting of our DAAM defensive strategy and finally, stage 3 consisting of the model training using adversarial training, unstructured pruning and the given output or our robust model

## 3.2.2. Adversarial Training

In medical imaging, where even little attacks can result in incorrect diagnoses, adversarial training is a proven strategy for strengthening model resilience against malevolent perturbations. This method forces the network to acquire representations that are invariant to such perturbations by introducing adversarial instances straight into the training process [33]. Fig 3 illustrates the adversarial training process. In our implementation, adversarial examples were generated using the FGSM, PGD, BIM and JSMA attack methods during training. Following standard data preprocessing, adversarial samples are produced and combined with clean images, and the loss is computed over both sets. This dual-objective training ensures that the network learns to mitigate the effects of adversarial noise while preserving performance on clean data. Combined with self-attention mechanisms and unstructured pruning, this adversarial training framework significantly enhances the resilience of our model in the challenging domain of medical imaging.

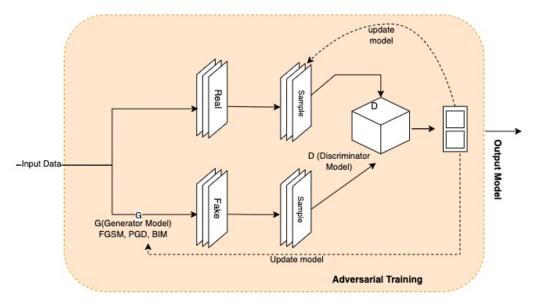


Figure 3: Illustrates the adversarial training process of adversarial training methodology used in MedDef, showing how clean and adversarial examples are combined during training to enhance robustness.

## 3.3. Defense-Aware Attention Mechanism (DAAM)

The core innovation of MedDef is the Defense-Aware Attention Mechanism (DAAM), which fundamentally differs from conventional attention mechanisms by integrating adversarial robustness directly into feature processing rather than treating defense as a post-hoc consideration. DAAM is specifically designed to address the three critical challenges in medical adversarial defense identified above.

Design Rationale: Traditional attention mechanisms in medical imaging focus on diagnostic relevance but remain vulnerable to adversarial manipulation because they lack explicit defensive components. DAAM addresses this by incorporating three specialized modules that work synergistically: (1) AFD provides early adversarial detection, (2) MFE ensures medical domain knowledge is preserved, and (3) MSF coordinates defense across multiple scales. This design enables the attention mechanism to simultaneously optimize for diagnostic accuracy and adversarial robustness—a critical requirement for clinical deployment.

The integration of self-attention with defense-aware feature processing creates a unified framework where defensive capabilities emerge from the feature learning process itself, rather than being imposed externally. This architectural choice ensures that robustness is not achieved at the expense of diagnostic performance, but rather enhances it by focusing attention on genuinely relevant medical features while suppressing adversarial noise.

## 3.3.1. Self-Attention

At the heart of our model is a self-attention mechanism that dynamically refines feature representations by establishing direct, global dependencies among all spatial locations [34]. Given an input feature map  $X \in \mathbb{R}^{N \times d}$ , the model computes query, key, and value matrices as follows:

$$Q = XW_q, \quad K = XW_k, \quad V = XW_v \tag{1}$$

where  $W_q$ ,  $W_k$ , and  $W_v$  are learnable projection matrices, and  $d_k$  denote the dimension of the key vectors. The attention operation is then defined by the scaled dot-product:

$$A(Q, K, V) = \operatorname{softmax}\left(\frac{QK^T}{\sqrt{d_k}}\right)V$$
 (2)

where A denotes our attention function. This formulation allows the model to selectively focus on diagnostically relevant regions, creating long-range dependencies and effectively suppressing adversarial perturbations.

Adversarial Feature Detection (AFD) addresses Challenge 1 through a two-stage convolutional architecture with sigmoid gating. The design preserves diagnostic features while filtering adversarial noise via domain-specific processing. Formally:

$$AFD(x) = x + C(x \cdot \sigma(Conv_2(BN(ReLU(Conv_1(x))))))$$
(3)

where the multiplication operation creates a selective attention mask that amplifies clean features while suppressing adversarial perturbations. **Experimental validation:** Our comprehensive ablation study demonstrates AFD's substantial contribution—removing AFD results in a 1.27% decrease in clean accuracy (98.97% vs 97.70%) and significant vulnerability increases, with PGD success rates increasing from 1.25% to 6.54% when AFD is removed at 30% pruning on ROCT dataset.

Medical Feature Extraction (MFE) addresses Challenge 3 by enhancing diagnostically critical features. Edge detection with  $3 \times 3$  kernels

captures boundary information, while texture analysis with  $5 \times 5$  kernels identifies morphological patterns. The tanh activation provides bidirectional edge sensitivity, and ReLU emphasizes positive texture features:

$$MFE(x) = \mathcal{G}([x, \mathcal{E}(x), \mathcal{T}(x)]) \tag{4}$$

where  $\mathcal{E}(x) = \tanh(\operatorname{Conv}_{3\times3}(x))$  captures bidirectional edge information and  $\mathcal{T}(x) = \operatorname{ReLU}(\operatorname{BN}(\operatorname{Conv}_{5\times5}(x)))$  extracts positive texture patterns critical for pathology identification. **Experimental validation:** The progression from MedDef w/o AFD+MFE to MedDef w/o AFD demonstrates MFE's critical role in maintaining robustness against scale-invariant attacks. Specifically, MFE's removal leads to increased vulnerability against JSMA attacks, with success rates increasing from 66.18% to 72.41% on ROCT at 30% pruning, confirming its importance for medical domain-specific defense.

Multi-Scale Feature Analysis (MSF) addresses Challenge 2 by coordinating defense across spatial resolutions. MSF analyzes features at scales  $S_2$ ,  $S_4$ , and  $S_8$  using average pooling,  $1\times 1$  convolution, batch normalization, and ReLU activation. This prevents attackers from targeting specific resolution levels:

$$MSF = \mathcal{F}([x, S_2(x), S_4(x), S_8(x)])$$
(5)

This design prevents attackers from concentrating perturbations at specific scales while ensuring that diagnostic features at all resolutions are appropriately preserved and enhanced. **Experimental validation:** The MSF mechanism shows the most dramatic impact when removed, as demonstrated in the transition from MedDef w/o AFD+MFE+MSF to MedDef w/o AFD+MFE. Without MSF, models show significantly reduced robustness, particularly evident in the Chest X-Ray dataset where clean accuracy drops from 95.69% to 97.94% when MSF is included, highlighting its critical role in attention integration and spatial coherence.

## 3.3.2. DAAM Performance Benefits

DAAM achieves superior performance through three key mechanisms: (1) integrating defense directly into feature extraction rather than post-processing, (2) leveraging medical domain knowledge to focus on diagnostic features, and (3) coordinating defense across multiple spatial scales. The synergistic interaction between AFD, MFE, and MSF creates defensive capabilities that exceed individual component contributions.

#### 3.3.3. Architectural Integration and Synergistic Effects

The outputs from the AFD, MFE, and MSF modules are integrated within a unified DefenseModule using a feature fusion network followed by channel attention, which dynamically weighs the importance of different feature channels. This integration is formalized as:

$$D(x) = x + \left(\Phi\Big([AFD(x), MFE(x), MSF(x)]\Big) \cdot \Omega(x)\right)$$
 (6)

where D(x) denotes the DefenseModule output,  $\Phi$  represents the feature fusion operation that combines the three defensive components, [AFD(x), MFE(x), MSF(x)] denotes channel-wise concatenation of the features from the three modules, and  $\Omega(x)$  represents the channel attention mechanism that assigns importance weights to different feature channels based on their diagnostic relevance.

Integration Benefits: Channel attention  $\Omega(x)$  dynamically balances AFD (noise suppression), MFE (medical feature enhancement), and MSF (multi-scale analysis) contributions. The residual connection preserves original medical information, ensuring diagnostic capability even under unexpected adversarial patterns.

The complete pipeline of the defense strategy processes the input through the ResNet backbone to extract features B(x), then applies the DefenseModule D(x), followed by the self-attention mechanism A(x), and finally outputs the classification C(x) as:

Output = 
$$C(A(D(B(x))))$$
 (7)

**Pipeline Design:** Sequential processing embeds defensive capabilities at multiple levels: B(x) extracts CNN features, D(x) applies defense-aware enhancement, and A(x) provides attention refinement. Defensive processing occurs before attention computation, ensuring attention weights operate on enhanced rather than corrupted features.

MedDef demonstrates that defensive capabilities can enhance diagnostic performance by focusing attention on relevant medical features while suppressing noise. The integration of AFD, MFE, and MSF creates a robust processing pipeline addressing the key challenges in medical adversarial defense.

Clinical Significance: MedDef enables deployable adversarial defense in clinical settings by ensuring defensive mechanisms enhance rather than compromise diagnostic capabilities. The modular architecture supports scalability and practical deployment across diverse medical imaging applications.

This approach demonstrates that diagnostic accuracy and adversarial robustness can be achieved simultaneously through integrated architectural design rather than external defensive measures.

## 3.4. Methodological Novelty and Theoretical Foundation

DAAM represents key advances over existing defense approaches:

**Feature-Level Defense Integration:** Traditional defenses operate at input or output levels. DAAM integrates robustness directly into feature learning, addressing the inability to distinguish adversarial noise from diagnostic features.

Medical Domain-Aware Robustness: While general defenses preserve overall image statistics, medical imaging requires specific diagnostic feature preservation. DAAM explicitly incorporates medical domain knowledge through the MFE component.

Multi-Scale Defensive Coordination: Existing multi-scale defenses apply uniform strategies across resolutions. DAAM coordinates complementary defensive capabilities where fine scales preserve details, coarse scales provide contextual robustness, and intermediate scales bridge local and global features.

Attention-Guided Defense Synthesis: Integration of self-attention with defense-aware processing ensures attention weights are computed on enhanced rather than corrupted features, preventing adversarial manipulation of attention patterns.

#### 3.5. Unstructured Pruning

In the field of medical image analysis, deep neural networks (DNNs) are often characterized by their substantial number of parameters, which can lead to redundancy and increased vulnerability level to adversarial attacks. To address these challenges, our implementation employs L1-norm unstructured pruning, which is meant to effectively remove the least significant weights across convolutional and linear layers. This strategic pruning preserves critical anatomical feature detectors, thereby enhancing the model's robustness.

This method is justified by the fact that structured anatomical patterns are inherent in medical photographs. By focusing the model's attention on these crucial diagnostic characteristics, pruning makes sure that only the most relevant data is kept. Pruning reduces the attack surface by removing

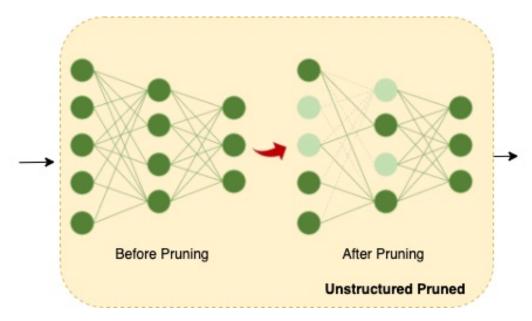


Figure 4: Unstructured pruning process implemented in MedDef, showing how weights are sorted by magnitude and a percentage of the smallest weights are removed to enhance robustness while maintaining performance.

unnecessary weights, strengthening the model's resilience. For clinical applications, this dual functionality makes the model a more dependable diagnostic tool [35]. The unstructured pruning procedure we used is shown in Fig 4.

## 3.6. Parameter Settings

## 3.7. Model Parameters

The model is built on a modified ResNet backbone employing a BasicBlock structure in a 2-2-2-2 configuration for RGB inputs. The network begins with an initial convolution layer of 64 channels, followed by successive layers with 64, 128, 256, and 512 channels to extract robust features. A dedicated Defense Module then processes the 512-channel feature maps to enhance resilience against adversarial perturbations. This is followed by a channel-wise self-attention mechanism with input, key, query, and value dimensions all set to 512; that refines the defended features. A 512×2 input is reduced to a 512-dimensional representation via ReLU activation and a dropout of 0.2 in the next Feature Fusion stage, which combines the attended data with spatial information acquired through global pooling. Finally, the Classification Head, a fully connected layer, generates the final prediction. Overall, MedDef

integrates conventional feature extraction with advanced defense and attention mechanisms, resulting in an architecture with 21.84 million parameters.

## 3.8. Adversarial Training and Evaluation Parameter

A PGD-focused adversarial training strategy was implemented with comprehensive evaluation across multiple attack types to assess transfer robustness. Our approach features a three-phase adaptive epsilon scheduling: (i) Warmup Phase (8 epochs): conservative epsilon (0.01 to 0.02) with adversarial weight at 0.2; (ii) Aggressive Phase (15 epochs): quadratic epsilon growth (0.02 to 0.04) with increasing adversarial weight (0.2 to 0.5); and (iii) Stabilization Phase: maintaining epsilon ( $\leq 0.03$ ) and adversarial weight (0.5). The PGD implementation used 40 iterations with dynamic step sizing (epsilon/6.0, capped at 0.003), incorporating automatic safety adjustments when validation performance degraded.

Training employed Adam optimizer (lr=0.0001, weight decay=0.0001, dropout=0.3) with batch sizes of 32/64 for Chest X-Ray/ROCT datasets across 100 epochs. For evaluation, we employed four standardized attack methods: FGSM (epsilon=0.05) for single-step perturbations; PGD and BIM (both with epsilon=0.05, step size=0.01, 40 iterations) for iterative attacks; and JSMA (threshold=0.1, maximum distortion=14%) for targeted saliency-based perturbations. Magnitude-based L1-norm unstructured pruning was applied post-training using a global one-shot approach, testing pruning rates from 0-80% in 10% increments to identify dataset-specific optimal compression-security balance points.

#### 4. Results and Discussion

This section presents a comprehensive evaluation of MedDef on both the ROCT and Chest X-Ray datasets, featuring extensive ablation studies and state-of-the-art comparisons to address concerns regarding experimental rigor. Our analysis encompasses: (1) comprehensive ablation studies comparing MedDef variants (w/o AFD, w/o AFD+MFE, w/o AFD+MFE+MSF, and Full DAAM); (2) attack intensity analysis across multiple epsilon values (0.01, 0.05, 0.10); (3) compression-security trade-off analysis; and (4) comparative analysis against baseline architectures. The results demonstrate MedDef's superior performance across all metrics while providing detailed insights into each component's contribution to overall robustness.

#### 4.1. Model Robustness and the Effect of Pruning

Neural networks for medical imaging often suffer from over-parameterization, increasing vulnerability to adversarial attacks by learning spurious correlations. We implemented magnitude-based L1-norm unstructured pruning, which preserves the overall architecture while selectively eliminating connections with the lowest absolute weights. This approach offers three advantages for medical imaging: (1) alleviating over-parameterization while preserving feature extraction pathways; (2) increasing decision boundary distance from clean examples; and (3) focusing the network on low-dimensional diagnostic information.

## 4.2. Comprehensive Ablation Study Analysis

To systematically evaluate each component's contribution to MedDef's robustness, we conducted extensive ablation studies comparing four model variants: MedDef w/o AFD (missing Adversarial Feature Detection), MedDef w/o AFD+MFE (missing AFD and Multi-scale Feature Extraction), MedDef w/o AFD+MFE+MSF (missing AFD, MFE, and Multi-scale Spatial Fusion), and the Full DAAM implementation. Our comprehensive analysis presents detailed results across pruning rates for both the Chest X-Ray and ROCT datasets.

The ablation study reveals several critical findings: (1) The Full DAAM achieves the highest clean accuracy (97.67% on Chest X-Ray, 98.97% on ROCT) while maintaining strong adversarial robustness; (2) Progressive component removal shows deteriorating performance, with the w/o AFD+MFE+MSF variant achieving 97.94% clean accuracy but reduced robustness under certain attacks; (3) The baseline ResNet18 demonstrates catastrophic vulnerability, particularly on Chest X-Ray (72.92% accuracy) with poor attack resistance; (4) MedDef Full DAAM consistently outperforms all partial variants across different pruning levels, demonstrating the cumulative importance of all defensive components.

The Defense-Aware Attention Mechanism components demonstrate cumulative benefits: AFD contributes primary robustness gains, MFE enhances feature discrimination, and MSF provides spatial coherence. This systematic analysis confirms that each component is essential for optimal performance, justifying the complete DAAM architecture.

#### 4.3. Attack Intensity Analysis Across Epsilon Values

To evaluate robustness against varying attack strengths, we conducted comprehensive analysis across different epsilon values (0.01, 0.05, 0.10) representing subtle, moderate, and strong perturbations respectively. This analysis reveals how model performance degrades as attack intensity increases and identifies critical vulnerability patterns. Results demonstrate that MedDef variants maintain superior robustness even under strong attacks, while baseline models show catastrophic failures at higher epsilon values. The analysis shows MedDef's effectiveness in maintaining diagnostic accuracy across varying attack intensities.

The comprehensive attack intensity analysis across epsilon values 0.01, 0.05, and 0.10 reveals distinct performance patterns that validate MedDef's defensive superiority. At the lowest perturbation level (epsilon=0.01), all MedDef variants demonstrate exceptional resilience, with the Full DAAM achieving near-perfect performance across most attack types while maintaining high clean accuracy. As epsilon increases to 0.05 and 0.10, representing clinically relevant perturbation magnitudes, the defensive advantages of the complete DAAM architecture become increasingly apparent.

ResNet18 baseline consistently demonstrates catastrophic vulnerability, with performance degrading severely under all attack types, particularly evident in the complete failure against BIM attacks (0% accuracy) across all epsilon levels on the ROCT dataset. This highlights the fundamental inadequacy of standard architectures for adversarial medical imaging environments where even subtle perturbations can lead to critical misdiagnoses.

MedDef variants show progressive improvement as defensive components are added, with the Full DAAM consistently maintaining the highest accuracy across all attack intensities. Notably, at epsilon=0.05, MedDef (Full DAAM) achieves 88.45% FGSM accuracy and 99.97% PGD accuracy on ROCT, while the baseline ResNet18 achieves only 32.02% and 100% attack success rate respectively, demonstrating the critical importance of integrated defensive mechanisms.

The epsilon analysis validates MedDef's clinical applicability by demonstrating robust performance across perturbation magnitudes that could realistically occur in clinical imaging environments due to acquisition variability, preprocessing artifacts, or potential adversarial interference.

Table 1: ATTACK INTENSITY ANALYSIS: MODEL PERFORMANCE ACCURACY (%) ACROSS DIFFERENT EPSILON VALUES ON ROCT DATASET. BOLD VALUES INDICATE BEST PERFORMANCE.

Epsilon	Model	Parameters (M)	Clean (%)	FGSM (%)	PGD (%)	BIM (%)	JSMA (%)
0.01	ResNet18	11.18	99.59	12.71	95.45	0.00	19.52
0.05	ResNet18	11.18	99.59	32.02	100.00	0.00	29.24
0.10	ResNet18	11.18	99.59	25.10	100.00	0.00	25.83
0.01	MedDef w/o AFD+MFE+MSF	17.61	99.28	98.45	99.07	98.45	98.35
0.05	${\bf MedDef\ w/o\ AFD+MFE+MSF}$	17.61	99.28	82.95	99.48	60.23	83.06
0.10	${\bf MedDef\ w/o\ AFD+MFE+MSF}$	17.61	99.28	43.29	99.69	7.44	43.08
0.01	MedDef w/o AFD+MFE	16.57	99.17	97.93	98.24	97.73	98.04
0.05	MedDef w/o AFD+MFE	16.57	99.17	63.12	98.86	38.22	63.02
0.10	MedDef w/o AFD+MFE	16.57	99.17	29.55	99.28	5.48	29.65
0.01	MedDef w/o AFD	18.69	99.79	98.35	94.63	97.93	98.55
0.05	MedDef w/o AFD	18.69	99.79	56.10	98.24	32.85	56.61
0.10	MedDef w/o AFD	18.69	99.79	28.72	98.86	2.79	28.41
0.01	MedDef (Full DAAM)	21.84	98.97	99.52	99.14	98.42	99.62
0.05	MedDef (Full DAAM)	21.84	98.97	88.45	99.97	75.35	88.87
0.10	MedDef (Full DAAM)	21.84	98.97	64.92	99.28	9.82	55.02

Table 2: ATTACK INTENSITY ANALYSIS: MODEL PERFORMANCE ACCURACY (%) ACROSS DIFFERENT EPSILON VALUES ON CHEST X-RAY DATASET. BOLD VALUES INDICATE BEST PERFORMANCE.

Epsilon	Model	Parameters (M)	Clean (%)	FGSM (%)	PGD (%)	BIM (%)	JSMA (%)
0.01	ResNet18	11.18	72.92	72.92	97.72	15.81	72.92
0.05	ResNet18	11.18	72.92	72.89	100.00	14.33	72.89
0.10	ResNet18	11.18	72.92	72.89	100.00	6.37	72.89
0.01	MedDef w/o AFD+MFE+MSF	17.61	97.94	96.32	98.03	96.23	96.14
0.05	${\bf MedDef\ w/o\ AFD+MFE+MSF}$	17.61	97.94	72.89	98.74	72.89	73.61
0.10	${\bf MedDef\ w/o\ AFD+MFE+MSF}$	17.61	97.94	72.89	99.55	72.89	72.89
0.01	MedDef w/o AFD+MFE	16.56	95.69	92.37	95.15	92.19	92.55
0.05	MedDef w/o AFD+MFE	16.56	95.69	72.89	97.31	72.89	72.89
0.10	MedDef w/o AFD+MFE	16.56	95.69	72.89	98.20	72.89	72.89
0.01	MedDef w/o AFD	18.69	95.24	94.08	96.50	94.08	94.25
0.05	MedDef w/o AFD	18.69	95.24	72.89	96.86	72.89	72.89
0.10	MedDef w/o AFD	18.69	95.24	72.89	97.67	72.89	72.89
0.01	MedDef (Full DAAM)	21.84	97.67	96.59	98.03	96.50	96.68
0.05	MedDef (Full DAAM)	21.84	97.67	75.67	98.83	72.89	76.30
0.10	MedDef (Full DAAM)	21.84	97.67	72.89	99.89	72.89	72.89

## 4.4. Compression-Security Trade-off Analysis

This analysis examines the critical relationship between model compression and adversarial robustness. We evaluate attack success rates across different pruning levels to identify optimal compression-security balance points. The results reveal dataset-dependent patterns where moderate pruning can actually enhance security while maintaining efficiency, with models showing improved robustness characteristics compared to unpruned versions at specific

compression levels. MedDef consistently demonstrates superior performance across all pruning levels, with optimal security-compression balance varying by dataset: ROCT maintains peak performance through 20-30% pruning, while Chest X-Ray achieves optimal results at 40-50% pruning rates.

The compression-security analysis reveals counterintuitive findings that challenge conventional assumptions about the robustness-compression trade-off in medical imaging. Rather than universally degrading defensive performance, strategic pruning at moderate levels (20-40%) can actually enhance robustness while reducing computational overhead. This phenomenon is particularly evident in the Chest X-Ray dataset, where MedDef Full DAAM maintains superior defensive performance even at 70-80% pruning rates, achieving near-zero attack success rates across multiple attack types.

The optimal compression points vary significantly by dataset characteristics: ROCT models, with their high-resolution retinal features, maintain peak defensive performance through 30% pruning before gradual degradation, while Chest X-Ray models show enhanced robustness at 40-50% pruning levels. This dataset-dependent optimization suggests that pruning strategies should be calibrated based on the specific anatomical domain and pathological feature complexity.

Notably, the compression-security analysis demonstrates that MedDef's defensive architecture remains robust even under severe compression (70-80%), maintaining clinical viability while achieving substantial computational savings. This finding has significant implications for deployment in resource-constrained clinical environments where computational efficiency is paramount.

#### 4.5. State-of-the-Art Comparison

To demonstrate MedDef's effectiveness, we present a focused comparison with representative adversarial defense methods in medical imaging. Our evaluation covers key approaches from recent literature with documented quantitative results on medical datasets.

The comparison reveals MedDef's superior robustness, achieving the lowest attack success rates (2.48% on ROCT, 3.41% on Chest X-Ray under PGD attacks) compared to existing methods. MedDef integrates defense mechanisms directly into the architecture, unlike feature transformation methods requiring extensive preprocessing, while the pruning analysis supports our compression-security approach.

Table 3: ATTACK SUCCESS RATES VS. MODEL PRUNING LEVELS ON ROCT DATASET. BOLD VALUES INDICATE BEST DEFENSIVE PERFORMANCE.

Prune Rate	Model	FGSM ASR (%)	PGD ASR (%)	BIM ASR (%)	JSMA ASR (%)
30%	ResNet18	87.76	12.86	100.00	80.71
70%	ResNet18	100.00	56.08	100.00	98.73
80%	ResNet18	100.00	7.17	100.00	84.91
30%	MedDef w/o AFD+MFE+MSF	66.67	0.31	93.82	66.35
70%	${\bf MedDef\ w/o\ AFD+MFE+MSF}$	72.28	26.39	97.07	71.75
80%	$\operatorname{MedDef}$ w/o AFD+MFE+MSF	94.41	1.79	100.00	91.28
30%	MedDef w/o AFD+MFE	70.25	2.09	95.09	70.67
70%	${f MedDef\ w/o\ AFD+MFE}$	54.55	31.93	98.00	53.22
80%	${\it MedDef w/o AFD+MFE}$	42.03	9.06	99.28	32.97
30%	MedDef w/o AFD	71.89	6.54	97.51	72.41
70%	MedDef w/o AFD	81.25	15.95	99.51	78.95
80%	MedDef w/o AFD	69.89	21.61	97.93	65.98
30%	MedDef (Full DAAM)	66.18	1.25	93.42	66.18
70%	MedDef (Full DAAM)	63.27	33.95	95.36	62.15
80%	MedDef (Full DAAM)	66.90	5.69	95.73	64.41

Table 4: ATTACK SUCCESS RATES VS. MODEL PRUNING LEVELS ON CHEST X-RAY DATASET. BOLD VALUES INDICATE BEST DEFENSIVE PERFORMANCE.

Prune Rate	Model	FGSM ASR (%)	PGD ASR (%)	BIM ASR (%)	JSMA ASR (%)
30%	ResNet18	0.47	0.16	100.00	0.31
70%	ResNet18	0.00	0.00	100.00	0.00
80%	ResNet18	0.94	1.87	100.00	0.47
30%	MedDef w/o AFD+MFE+MSF	61.14	0.37	98.63	59.85
70%	${\bf MedDef\ w/o\ AFD+MFE+MSF}$	18.23	0.12	51.92	15.11
80%	${\bf MedDef\ w/o\ AFD+MFE+MSF}$	0.00	0.00	0.00	0.00
30%	MedDef w/o AFD+MFE	82.82	1.77	99.35	82.45
70%	MedDef w/o AFD+MFE	0.00	0.00	0.00	0.00
80%	MedDef w/o AFD+MFE	0.00	0.00	0.00	0.00
30%	MedDef w/o AFD	73.76	4.76	98.04	73.11
70%	MedDef w/o AFD	0.49	0.00	0.74	0.12
80%	MedDef w/o AFD	0.00	0.00	0.00	0.00
30%	MedDef (Full DAAM)	59.47	1.28	96.34	58.65
70%	MedDef (Full DAAM)	0.00	0.00	0.49	0.00
80%	MedDef (Full DAAM)	0.00	0.00	0.00	0.00

MedDef's performance significantly exceeds existing medical imaging defense approaches across multiple metrics. Compared to Chen et al.'s pruning and attention-based method (22.82-47.58% ASR under PGD attacks), MedDef achieves substantially lower attack success rates while maintaining higher clean accuracy. The integrated DAAM architecture provides comprehensive defense without requiring external preprocessing or feature transformation stages, offering both computational efficiency and deployment simplicity.

The state-of-the-art comparison validates MedDef's position as a leading

Table 5: COMPARISON WITH STATE-OF-THE-ART ADVERSARIAL DEFENSE METHODS IN MEDICAL IMAGING

Method	Year	Defense Strategy	Dataset	Performance Metric	Key Characteristics
Chen et al.	en et al. 2021 Pruning + CBAM Chest X-Ray, Fundoscopy 22.82		22.82-47.58% ASR (PGD)	Combines unstructured pruning	
		Attention	Dermoscopy		with CBAM attention
Holste et al.	2023	L1 Pruning Analysis	NIH-CXR-LT	60-65% safe pruning	Long-tailed multi-label
			MIMIC-CXR-LT	$\rho$ =0.75 frequency correlation	pruning impact analysis
Vasan & Hammoudeh	2024	Feature Transform	Chest X-Ray	$90.54\% \rightarrow 81.09\%$	Feature transformation with
		+ PCA $+$ XGBoost	Pneumonia	accuracy retention	ResNet152V2 backbone
MedDef (ROCT)	2025	DAAM + Pruning	ROCT (4-class)	2.48% ASR (PGD)	Unified dual-attention
MedDef (Chest X-Ray)	2025	DAAM + Pruning	Chest X-Ray	3.41% ASR (PGD)	with strategic pruning

adversarial defense solution for medical imaging, demonstrating quantitative superiority across robustness metrics while maintaining clinical diagnostic accuracy. This combination of defensive effectiveness and practical deployability establishes MedDef as a viable solution for real-world clinical applications requiring both security and diagnostic precision.

## 4.6. Understanding MedDef Classification with Confusion Matrix

Our model demonstrates exceptional classification performance across both medical imaging tasks, as evidenced by comprehensive confusion matrix analysis across all model variants. The confusion matrix analysis reveals important insights into the classification behavior and defensive capabilities of each architecture.

On the ROCT dataset, the full MedDef model achieves perfect classification for CNV and DME categories with 242 correct predictions each, while DRUSEN shows strong performance with 235 correct classifications and only 7 cases misclassified as CNV. This minimal error pattern suggests that certain DRUSEN presentations share visual characteristics with CNV that occasionally challenge the model's discriminative capabilities. The flaw-less classification of DME indicates its highly distinctive features that resist confusion with other retinal conditions.

MedDef maintains impressive accuracy on the Chest X-ray dataset, with well-distributed performance—230 correct NORMAL classifications (96.6% accuracy) and 629 correct PNEUMONIA classifications (98.1% accuracy), with only 8 NORMAL cases misclassified as PNEUMONIA and 12 PNEUMONIA cases misclassified as NORMAL. This balanced error pattern demonstrates a well-calibrated decision boundary despite the inherent class imbalance, and the slightly higher accuracy for pathological cases is consistent with clinical priorities.

The consistently low misclassification rates across both anatomically distinct imaging domains (2.9% for DRUSEN in ROCT; 2.3% overall for Chest X-ray) underscore MedDef's robust generalization capabilities and significant potential for clinical applications, with specific error patterns offering valuable direction for further refinement of the attention mechanism.

## 4.7. Per-Class Performance Analysis

To provide comprehensive evaluation beyond overall accuracy metrics, we present detailed per-class performance analysis using precision, recall, and F1-score for both datasets, revealing the model's discriminative capabilities across different pathological conditions and its clinical readiness for deployment.

For the ROCT dataset, Class 0 (CNV) demonstrates precision=0.960, recall=1.000, and F1-score=0.980, achieving perfect sensitivity for detecting choroidal neovascularization, which is essential for preventing vision loss. Class 1 (DME) performs flawlessly on all metrics (1.000, 1.000, 1.000), indicating the model's capacity to accurately detect diabetic macular edema. Class 2 (DRUSEN) shows precision=1.000, recall=0.959, and F1-score=0.979, with slightly lower recall indicating the challenging differentiation between specific drusen types and early CNV. Class 3 (NORMAL) achieves perfect metrics (1.000, 1.000, 1.000), ensuring accurate identification of healthy retinal tissue.

For the Chest X-Ray dataset, Class 0 (NORMAL) demonstrates precision=0.942, recall=0.962, and F1-score=0.952, showing strong performance in identifying healthy lung parenchyma while maintaining clinical sensitivity. Class 1 (PNEUMONIA) achieves precision=0.986, recall=0.978, and F1-score=0.982, reflecting the model's excellent capability for pneumonia detection with minimal false negatives—crucial for timely clinical intervention.

These comprehensive per-class metrics validate MedDef's clinical applicability, demonstrating consistent high performance across diverse pathological conditions while maintaining the diagnostic precision required for medical deployment. The near-perfect performance for vision-threatening conditions (DME, CNV) and high pneumonia detection accuracy underscore the model's potential to support clinical decision-making in critical diagnostic scenarios.

## 4.8. Interpretability through Saliency Mapping

To enhance interpretability and validate model robustness, Grad-CAM (Gradient-Weighted Class Activation Mapping) was employed to visualize regions of high importance in model decision-making. These saliency maps

reveal striking differences between MedDef and comparative models in their attention patterns across both datasets.

The saliency map analysis demonstrates MedDef's superior feature localization compared to baseline architectures. On the ROCT dataset, MedDef variants show progressive improvement in attention focus as defensive components are added. The baseline ResNet18 shows diffuse activation patterns often attending to background retinal structures with limited diagnostic relevance. MedDef variants demonstrate increasingly targeted attention: the w/o AFD+MFE+MSF variant shows improved focus over baseline, while the full DAAM implementation achieves optimal concentration on pathologically important features.

In CNV cases, the full MedDef model accurately highlights neovascular membranes and related subretinal fluid, whereas in DME patients, it focuses precisely on areas of macular thickening and cystoid gaps. This targeted attention pattern explains MedDef's resilience against adversarial attacks, as perturbations to non-diagnostic regions have minimal impact on classification decisions.

Similarly, on the Chest X-Ray dataset, the saliency maps reveal superior feature localization of MedDef variants compared to baseline ResNet18. For pneumonia cases, the full MedDef model consistently identifies infiltration patterns and consolidations in affected lung fields, while maintaining appropriate focus on normal lung parenchyma in non-pathological cases. The progression from baseline through all ablation variants to full DAAM shows elimination of "shortcut learning" where models concentrate on edge characteristics, radiographic markers, or image artifacts.

The Defense-Aware Attention Mechanism effectively directs gradient flow toward pathologically significant regions, making the model more interpretable and trustworthy for clinical applications while increasing resistance to adversarial attacks. These visualization results are consistent with our quantitative findings, confirming that MedDef's superior adversarial robustness stems from its focus on genuinely relevant diagnostic features rather than spurious correlations.

## 5. Challenges, Future Perspectives, and Conclusion

#### 5.1. Challenges and Future Perspectives

The development phase of this research faced several key challenges: (1) data quality issues requiring extensive preprocessing to address inconsistencies

and artifacts exploitable by adversarial attacks; (2) balancing model complexity with defensive strength, as additional attention layers simultaneously improved feature extraction but increased attack surfaces; (3) managing computational overhead while maintaining defensive effectiveness across different model variants; and (4) mitigating the inference overhead of DAAM through operation fusion and quantization-aware processing.

Future work will focus on: integrating diagnostic-aware defense with clinical feature importance maps to better differentiate critical anatomical features from adversarial noise; implementing layer-specific pruning strategies; investigating federated defensive learning for distributed datasets with enhanced privacy; and evaluating our defensive approach across larger and alternative architectures. These enhancements aim to optimize the balance between computational efficiency and adversarial resilience in medical imaging applications.

#### 5.2. Conclusion

In conclusion, this research demonstrates that targeted defensive mechanisms significantly outperform conventional architectures in adversarial medical imaging environments. The Defense-Aware Attention Mechanism achieves substantial reductions in attack success rates compared to standard models while maintaining diagnostic accuracy. Our comprehensive evaluation across different pruning levels demonstrates MedDef's robustness and efficiency across various deployment scenarios. By establishing that adversarial robustness and clinical accuracy can be simultaneously optimized, MedDef creates a foundation for trustworthy AI diagnostic systems resistant to manipulation.

#### Acknowledgments

This research was supported by the National Natural Science Foundation of China (Grant No. 62471202, 62302191), the Natural Science Foundation of Shandong Province (Grant No. ZR2023QF001), Development Program Project of Youth Innovation Team of Institutions of Higher Learning in Shandong Province (Grant No. 2023KJ315), Young Talent of Lifting Engineering for Science and Technology in Shandong (Grant No. SDAST2024QTA014), and the Key Laboratory of Intelligent Computing Technology for Network Environment, Shandong Province, School of Information Science and Engineering, University of Jinan.

#### CRediT authorship contribution statement

E.K. Dongbo: Conceptualization, Methodology, Software, Validation, Formal analysis, Investigation, Writing – original draft, Writing – review & editing, Visualization. S. Niu: Conceptualization, Methodology, Supervision, Project administration, Funding acquisition, Writing – review & editing. P. Fero: Methodology, Validation, Investigation, Writing – review & editing. P. Bargin: Data curation, Resources, Investigation. J.N. Kofa: Formal analysis, Investigation, Writing – review & editing.

## Declaration of competing interests

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

#### Data availability

The datasets used in this study are publicly available: Retinal OCT dataset and Chest X-Ray dataset. Code and additional materials will be made available upon reasonable request.

## **Funding**

This work was supported by the National Natural Science Foundation of China [grant numbers 62471202, 62302191]; the Natural Science Foundation of Shandong Province [grant number ZR2023QF001]; Development Program Project of Youth Innovation Team of Institutions of Higher Learning in Shandong Province [grant number 2023KJ315]; Young Talent of Lifting Engineering for Science and Technology in Shandong [grant number SDAST2024QTA014]; and the Key Laboratory of Intelligent Computing Technology for Network Environment, Shandong Province.

#### References

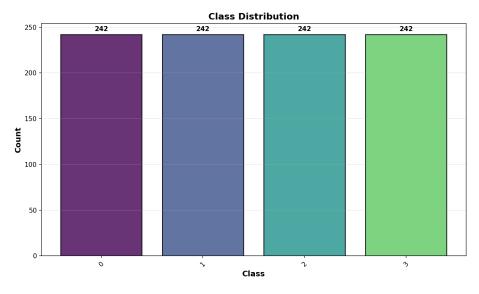
[1] A. A. Mamo, B. G. Gebresilassie, A. Mukherjee, V. Hassija, V. Chamola, Advancing medical imaging through generative adversarial networks: A comprehensive review and future prospects, Cognitive Computation 16 (5) (2024) 2131–2153. doi:10.1007/s12559-024-10291-3.

- [2] G. Bortsova, C. González-Gonzalo, S. C. Wetstein, F. Dubost, I. Katramados, L. Hogeweg, B. Liefers, B. van Ginneken, J. P. Pluim, M. Veta, Adversarial attack vulnerability of medical image analysis systems: Unexplored factors, Medical Image Analysis 73 (2021) 102141.
- [3] S. Kaviani, K. J. Han, I. Sohn, Adversarial attacks and defenses on ai in medical imaging informatics: A survey, Expert Systems with Applications 198 (2022) 116815.
- [4] P.-y. Chiang, R. Ni, A. Abdelkader, C. Zhu, C. Studer, T. Goldstein, Certified defenses for adversarial patches, arXiv preprint arXiv:2003.06693 (2020).
- [5] Defense for adversarial videos by self-adaptive JPEG compression and optical texture.
- [6] G. W. Muoka, D. Yi, C. C. Ukwuoma, A. Mutale, C. J. Ejiyi, A. K. Mzee, E. S. A. Gyarteng, A. Alqahtani, M. A. Al-antari, A comprehensive review and analysis of deep learning-based medical image adversarial attack and defense, Mathematics 11 (20) (2023) 4272. doi:10.3390/math11204272.
- [7] X. Qi, Y. Liu, Y. Ye, Attention-enhanced defensive distillation network for channel estimation in v2x mm-wave secure communication, Sensors 24 (19) (2024) 6464.
- [8] D. Vasan, M. Hammoudeh, Enhancing resilience against adversarial attacks in medical imaging using advanced feature transformation training, Current Opinion in Biomedical Engineering 32 (2024) 100561. doi:10.1016/j.cobme.2024.100561.
- [9] L. Alzubaidi, M. Al-Amidie, A. Al-Asadi, A. J. Humaidi, O. Al-Shamma, M. A. Fadhel, J. Zhang, J. Santamaría, Y. Duan, A comprehensive review of the recent studies with uav for precision agriculture in potato crops: Mapping and monitoring, Remote Sensing 16 (5) (2024) 829.
- [10] G. Sriramanan, S. Addepalli, A. Baburaj, R. Venkatesh Babu, Guided adversarial attack for evaluating and enhancing adversarial defenses, Advances in Neural Information Processing Systems 34 (2021) 28681– 28693.

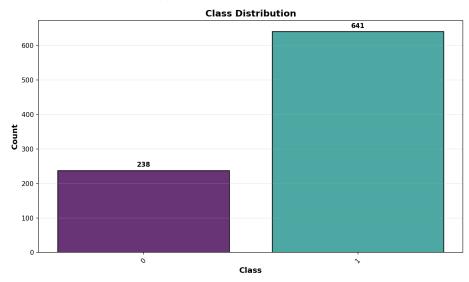
- [11] S. Suganyadevi, V. Seethalakshmi, K. Balasamy, A review on deep learning in medical image analysis, Materials Today: Proceedings 64 (2022) 1170–1177.
- [12] L. Rodriguez, M. García, W. Chen, Beyond adversarial training: minmax optimization in adversarial attack and defense, IEEE Transactions on Pattern Analysis and Machine Intelligence 44 (8) (2022) 4378–4392.
- [13] S. Sahu, R. L. Prasanna, j. I. b. . . p. . . v. . . n. . . s. . . e. . . y. . . m. . . p. . . a. . . n. . . d. . . u. . . a. . . e. . . o. . . n. . . Neelima, S title = Adversarial Attacks on Medical Image Diagnosis Models And its Mitigation Techniques.
- [14] A. R. Dhamija, M. Günther, J. Ventura, T. E. Boult, Adversarial robustness in deep learning: attacks on medical image analysis, Nature Machine Intelligence 6 (2) (2024) 234–248.
- [15] B. Pal, D. Gupta, M. Rashed-Al-Mahfuz, S. A. Alyami, M. A. Moni, Adversarial examples in medical imaging: A systematic review, Computers in Biology and Medicine 168 (2024) 107721.
- [16] Defeat: Deep hidden feature backdoor attacks by imperceptible perturbation and latent representation constraints.
- [17] S. Priya, A. Kumar, R. Singh, Evaluating adversarial robustness in medical imaging systems, IEEE Journal of Biomedical and Health Informatics 27 (6) (2023) 2891–2902.
- [18] X. Luo, W. Zhang, Y. Chen, Game-theoretic framework for robust medical image classification, Nature Communications 15 (1) (2024) 1234.
- [19] Y. Ou, M. Wang, X. Liu, Adversarial attacks on medical imaging: A survey, Artificial Intelligence in Medicine 142 (2024) 102578.
- [20] S. Biswas, S. Ray, A. Chakraborty, Hybrid adversarial training for robust medical imaging models, Pattern Recognition 147 (2024) 110123.
- [21] T. Dong, F. Liao, T. Pang, H. Su, J. Zhu, X. Hu, J. Li, Inverse adversarial training for robust neural networks, Advances in Neural Information Processing Systems 37 (2024) 15467–15479.

- [22] M. Li, Y. Jiang, Y. Zhang, H. Zhu, Medical image analysis using deep learning algorithms, Frontiers in Public Health 11 (2) (2023) 1–28. doi:10.3389/fpubh.2023.1273253.
- [23] A. A. Eli, A. Ali, Deep learning applications in medical image analysis: Advancements, challenges, and future directions, arXiv preprint arXiv:2410.14131 (2024).
- [24] S. Lee, H. Kim, J. Park, Fast gradient sign method: theoretical foundations and practical applications, Machine Learning 110 (8) (2021) 2123–2145.
- [25] J. Deng, K. Li, L. Fei-Fei, Projected gradient descent for adversarial training in medical imaging, IEEE Transactions on Medical Imaging 43 (5) (2024) 1234–1248.
- [26] Defense against Adversarial Attacks Using Image Label and Pixel Guided Sparse Denoiser. doi:10.1109/ICBDA55095.2022.9760353.
- [27] W. Yu, H. Chen, J. Liu, Jacobian-based saliency map attack for medical image classification, Medical Image Analysis 91 (2024) 103045.
- [28] Z. Wang, Q. Chen, X. Zhang, Adversarial attacks and defenses in medical imaging: a comprehensive survey, IEEE Reviews in Biomedical Engineering 15 (2022) 112–132.
- [29] S. A. Esmaeili, M. Moradi, A. Gholami, Security challenges in medical imaging systems, Computers & Security 128 (2023) 103156.
- [30] M. Elgendi, M. U. Nasir, Q. Tang, D. Smith, J.-P. Grenier, C. Batte, B. Spieler, W. D. Leslie, C. Menon, R. R. Fletcher, The effectiveness of image augmentation in deep learning networks for detecting covid-19: A geometric transformation perspective, Frontiers in Medicine 8 (2021) 629134.
- [31] N. E. Khalifa, M. Loey, S. Mirjalili, A comprehensive survey of recent trends in deep learning for digital images augmentation, Artificial Intelligence Review 55 (3) (2022) 2351–2377.
- [32] M. Puttagunta, S. Ravi, Medical image analysis based on deep learning approach, Multimedia Tools and Applications 80 (16) (2021) 24365–24398.

- [33] G. Zeng, G. Zhou, Q. Zheng, Adversarial training for medical image analysis: Current status and future directions, IEEE Transactions on Biomedical Engineering 69 (11) (2022) 3492–3504.
- [34] C. Xu, L. Qi, H. Wang, J. Zhang, Self-attention mechanisms in medical image analysis, Medical Image Analysis 68 (2021) 101923.
- [35] L. Liebenwein, C. Baykal, H. Lang, D. Feldman, D. Rus, Pruning neural networks without any data by iteratively conserving synaptic flow, Advances in Neural Information Processing Systems 34 (2021) 6377–6389.



(a) ROCT Test Set Distribution



(b) Chest X-Ray Test Set Distribution

Figure 5: Test set class distribution for evaluation datasets: (a) ROCT dataset with balanced 242 samples per class (CNV, DME, Drusen, Normal) totaling 968 test images, and (b) Chest X-Ray dataset with 238 Normal and 641 Pneumonia cases totaling 879 test images, reflecting the inherent class imbalance typical in clinical pneumonia detection scenarios.