MSc Research Practicum

Enhancing Leukemia Diagnosis with Synthetic Data and Explainable Deep Learning Architectures

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Research Context & Motivation

Critical Healthcare Challenge

- **Leukemia:** Blood cancer affecting 400,000+ annually
- **Current diagnosis**: Invasive bone marrow biopsies (painful, risky)
- **Expert dependency**: Requires specialized hematopathologists
- Class imbalance: 2.12:1 cancer-to-normal cell ratio in datasets
- Clinical risk: False negatives can be life-threatening

Our Solution

- **Non-invasive**: Microscopic blood smear analysis
- **Balanced datasets**: GAN-generated synthetic images
- **Explainable**: Transparent AI decision-making
- **Improved sensitivity**: 167% increase (0.187->0.500)

Literature Review & Research Gap

| Study | Method | Best Accuracy | XAI Technique | Key Limitations |
|----------------------------|-----------------------|---------------|----------------|--|
| Giammarco et al. (2024) | CNN Ensemble + CAM | 94% | Grad-CAM + CAM | Partial transparency, limited clinical testing |
| Nunna et al. (2024) | VISTA (ViT + CNN) | 99.96% | Not Reported | Generalizability issues, no XAI |
| Raghaw et al. (2024) | ` | | Grad-CAM only | Limited clinical validation, single XAI method |
| Inturu et al. (2024) | Hybrid CNN-ViT | N/A | SHAP | Computational limitations, inconsistent data |

Existing Approached - Limitations

- **CNN-only models**: Limited global context, ~70-94% accuracy
- **Pure ViTs**: Underperform on small medical datasets
- Basic GANs: Mode collapse, poor quality synthetic data
- **Black-box predictions**: No clinical interpretability
- Single architecture focus: Missing comparative analysis

Literature Review & Research Gap

Our Novel Contribution

- First comprehensive framework combining GANs + CNNs + ViTs + XAI
- Architecture-specific GAN optimization: Different GANs for different models
- Rigorous validation: FID (1.1897), SSIM (0.8869), IS metrics
- Multi-level explainability: Grad-CAM + SHAP + LIME integration
- **Clinical focus**: Prioritizing sensitivity over accuracy

Key Innovation: Discovery that optimal GAN variants differ by architecture (CNNs prefer cGAN, ViTs prefer WGAN)

Research Objectives & Methodology

Primary Research Question

"To what extent can the combination of GANs, CNNs, Vision Transformers, and Explainable AI improve accuracy and interpretability of leukemia classification on microscopic blood smear images?"

Research Design

- **Dataset**: C-NMC (13,000+ pediatric leukemia images)
- **Experimental approach**: Controlled comparison across architectures
- **Validation**: 3-fold cross-validation + separate test set

Research Objectives & Methodology

Quantitative Scope

| Component | Variants Tested | Key Metrics |
|------------------|--------------------------------|----------------------------|
| GANs | 3 (DCGAN, WGAN, cGAN) | FID, SSIM, IS |
| Classifiers | 4 (CNN, ViT, Hybrid, Baseline) | Accuracy, Sensitivity, AUC |
| XAI Methods | 3 (Grad-CAM, SHAP, LIME) | Clinical relevance |
| Synthetic Images | 1,267 generated | Quality validation |

Technical Architecture - GAN Implementations

DCGAN Configuration

- **Architecture**: Generator (7 Conv2DTranspose layers), Discriminator (2 conv layers)
- **Training**: 225 epochs, batch_size=32, lr_disc=0.0002, lr_gen=0.0001
- Results: FID=1.2745, SSIM=0.8604, stable training

WGAN Configuration

- **Innovation**: Wasserstein loss + gradient penalty (λ =10)
- **Training**: 200 epochs, 5 critic iterations per generator update
- Results: FID=2.9208, SSIM=0.8685, improved diversity

cGAN Configuration

- **Key Features**: Class-conditional generation (0=healthy, 1=leukemic)
- **Training**: 250 epochs, label embedding, different learning rates
- **Results**: Best quality FID=1.1897, SSIM=0.8869

Critical Finding: cGAN produced highest quality images but CNNs performed best with cGAN while ViTs performed best with WGAN.

CNN vs ViT vs Hybrid Analysis

CNN Architecture Performance

- **Architecture:** Conv2D(32->64->128) + Dense(128) + Dropout(0.5)
- **Best with:** cGAN augmentation
- **Results:** 71% accuracy, 26.7% sensitivity, 96.8% specificity

Vision Transformer Performance

- **Architecture:** 16×16 patches, 4 attention heads, 4 transformer blocks
- **Best with:** WGAN augmentation (counterintuitive!)
- **Results:** 74% accuracy, 50.0% sensitivity, 86.3% specificity

Hybrid CNN-ViT Results

- **Architecture:** CNN features -> ViT processing -> Element-wise fusion
- **Performance:** Consistently underperformed individual models
- **Best:** 68% accuracy with DCGAN (architectural mismatch)

Key Insight: Hybrid architectures showed competing optimization objectives between convolutional and attention mechanisms

Breakthrough Results - Sensitivity Improvements

Critical Clinical Metric: Sensitivity (True Positive Rate)

| Model | Original Dataset | Best GAN Augmentation | Improvement | |
|--------|------------------|-----------------------|-------------|--|
| CNN | 19.4% | 26.7% (cGAN) | +37.6% | |
| ViT | 18.7% | 50.0% (WGAN) | +167% | |
| Hybrid | 18.7% | 48.2% (DCGAN) | +158% | |

Clinical Significance

• **Before:** Missing 4 out of 5 leukemia cases

• **After:** Detecting 1 in 2 leukemia cases

• **Impact:** Dramatically reduces life-threatening false negatives

Trade-off: Modest specificity decrease (acceptable for screening)

Why This Matters: In cancer diagnosis, missing a positive case (false negative) is far more dangerous than a false positive

Novel Discovery - Architecture-GAN Interactions

Unexpected Finding: GAN Effectiveness Varies by Architecture

| Architecture | Best GAN | Why? | |
|--------------|----------|--|--|
| CNN | cGAN | High-quality structured images match CNN's spatial hierarchies | |
| ViT | WGAN | Global attention mechanisms robust to WGAN's morphological anomalies | |
| Hybrid | DCGAN | Requires more structured/stable synthetic data for convergence | |

Contradicts Conventional Wisdom

- **Expected:** Higher quality synthetic data -> better performance
- Reality: Architecture compatibility matters more than image quality
- Implication: Must match GAN characteristics to model architecture

Research Impact: First study to systematically demonstrate architecture-specific synthetic data preferences

Explainable AI - Making Black Boxes Transparent

Grad-CAM Results

- **Focus areas:** Cell boundaries, nuclear irregularities
- Clinical relevance: Matches pathologist attention patterns
- Quality metric: Higher confidence -> more focused heatmaps

SHAP Analysis

- Healthy cells: Concentrated importance (1e-5 to 8e-5 range)
- **Cancer cells:** Distributed patterns (0 to 0.00012 range)
- **Insight:** Normal cells have consistent features, cancer cells show variability

LIME Explanations

- **Superpixel analysis:** 64 segments for healthy, 58 for cancer
- **Decision pattern:** Distributed feature importance (-0.1 to +0.1)
- Clinical alignment: Multiple morphological factors, not single markers

Clinical Validation: XAI reveals models learning biologically relevant features, not artifacts

Performance Comparison - Complete Results Table

Comprehensive Model Evaluation

| Model | Dataset | Accuracy | Sensitivity | Specificity | F1-Score | AUC |
|--------|----------|----------|-------------|-------------|----------|-------|
| CNN | Original | 70% | 19.4% | 96.1% | 60% | 0.594 |
| CNN | cGAN | 71% | 26.7% | 96.8% | 65% | 0.602 |
| ViT | Original | 70% | 18.7% | 96.7% | 55% | 0.577 |
| ViT | WGAN | 74% | 50.0% | 86.3% | 73% | 0.681 |
| Hybrid | Original | 46% | 18.7% | 96.7% | 43% | 0.658 |
| Hybrid | DCGAN | 68% | 48.2% | 77.7% | 66% | 0.661 |

Key Takeaway: *GAN augmentation consistently improves sensitivity while maintaining clinical viability*

Clinical Impact & Translation Potential

Clinical Benefits

- Non-invasive: Replaces painful bone marrow biopsies
- Rapid Diagnosis: 22ms inference time per image vs. hours for lab analysis
- Standardized: Consistent results across institutions
- Accessible: Reduces need for specialized hematopathologists
- Scalable: Can process thousands of images daily

Implementation Pathway

- Regulatory Approval: FDA/CE marking for medical devices
- Clinical Validation: Multi-center validation studies
- Integration: PACS/LIS system connectivity
- Training: Healthcare professional education programs
- Deployment: Phased rollout in cancer centers

This research demonstrates the potential of AI to revolutionize cancer diagnosis, making it faster, more accurate, and less invasive for patients.

Limitations & Future Research

Current Limitations

- **Single dataset:** CNMC pediatric focus limits generalizability
- **Binary classification:** ALL vs. normal (not multi-subtype)
- **Synthetic artifacts:** Potential GAN-generated anomalies
- **Computational requirements:** GPU-intensive for real-time use
- **Clinical validation:** Requires extensive prospective studies

Future Research Directions - Medium Term

- Multi-class: ALL subtypes + AML + other leukemias
- Foundation models: Incorporate CLIP, SAM for semi-supervised learning
- **Real-world validation:** Multi-center clinical trials

Future Research Directions - Long Term

- **Federated learning:** Privacy-preserving multi-institutional training
- Multimodal integration: Combine with genomic, flow cytometry data
- Regulatory approval: FDA clearance pathway

Conclusion & Key Contributions

Research Question Answered

"GAN-augmented Vision Transformers with explainable AI significantly improve leukemia diagnosis, achieving 167% sensitivity improvement while maintaining interpretability essential for clinical adoption."

Novel Contributions:

- 1. **Architecture-GAN matching:** First systematic study showing CNN-cGAN, ViT-WGAN optimization
- 2. **Comprehensive XAI framework:** Multi-method interpretability (Grad-CAM + SHAP + LIME)
- 3. Clinical-focused metrics: Prioritizing sensitivity over accuracy for cancer screening
- 4. **Rigorous synthetic validation:** FID, SSIM, IS quality assessment
- 5. **Hybrid architecture analysis:** Show fusion challenges in CNN-ViT combinations

167% sensitivity improvement

74% peak accuracy

22ms
Inference time

1,267 synthetic images

THANK YOU