Histopathological Tissue Based Cancer Tumor Detection

Chenyu Wang, Hector Rincon, Ifrah Javed, Justin To and Srila Maiti

AGENDA

PROJECT

01

MOTIVATION

Research Question and Application DATASET AND

AUGMENTATION

02

Data Source and Feature Engineering

OUR APPROACH

03

Iterative Process, Metrics

Architecture and

Architecture and Performance

05

FINAL MODEL FUTURE WORK

Further experimentation and questions

01

PROJECT MOTIVATION

PROJECT BACKGROUND AND NEED

KEY CANCER TREATMENT FACTORS

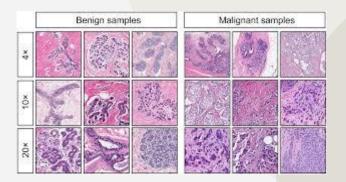
CORRECT EARLY
DIAGNOSIS INTERVENTION

DEATH RATE



609,360 Deaths out of 1.9 million cases in 2022

CURRENT DIAGNOSTIC METHOD



MISDIAGNOSIS RATE

24.7%

RESEARCH QUESTION

"CAN WE PREDICT IF A TUMOR IS
BENIGN OR MALIGNANT BASED ON A
HISTOPATHOLOGICAL TISSUE IMAGE?"

PROJECT APPLICATION

PATIENT PRESENTS SYMPTOMS

Start of care.



OUR ALGORITHM

A diagnosis is provided by running our algorithm.



BIOPSY

Surgical process to remove cells and create a histopathological image.



TREATMENT

Treatment plan is decided based on diagnosis.

02

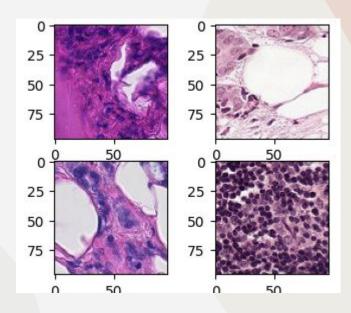
DATASET & AUGMENTATION

DATASET

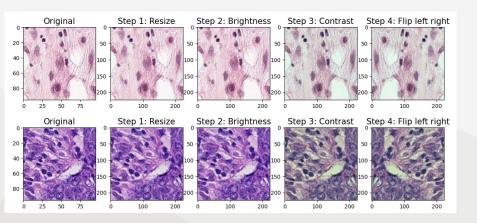
- Working with image data of pathological cells for cancer detection
- A version of the PCAM data set (https://github.com/basveeling/pcam)
- Cleaned from duplicates by Kaggle for the <u>Histopathologic</u>
 <u>Cancer Detection competition</u>
- Dataset contains 277,483 total images (220,025 train / 57,458 test)
- Test images are not labeled so we repurposed the 220K images and made the split on that subset
- Positive label: center 32x32px (out of 96x96 image) region of a patch contains at least one pixel of tumor tissue

[1] B. S. Veeling, J. Linmans, J. Winkens, T. Cohen, M. Welling. "Rotation Equivariant CNNs for Digital Pathology". arXiv:1806.03962

[2] Ehteshami Bejnordi et al. Diagnostic Assessment of Deep Learning Algorithms for Detection of Lymph Node Metastases in Women With Breast Cancer. JAMA: The Journal of the American Medical Association, 318(22), 2199–2210. doi:jama.2017.14585



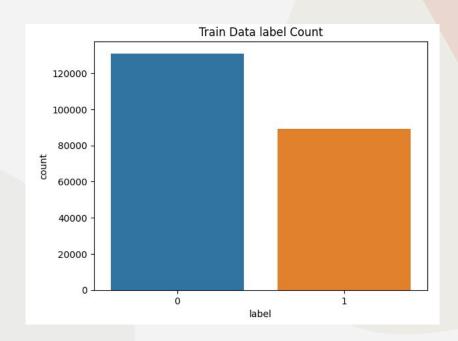
DATA AUGMENTATION



- Converted to grayscale (no info from color data)
- Attempted to use ImageDataGenerator (does not add images, but returns the new transformations)
- Aug. steps:
 - Contrast
 - Brightness
 - Random flips
 - Rotation (90, 180, 270) comparison
- Attempted to crop to 32x32/64x64 center region
- Limitations/calls of judgement

UNBALANCED DATA

- The training dataset had an imbalance of the positive vs negative examples (negative overrepresented).
- Solution: undersample the negative examples (took a random sample of # of positive examples available in the dataset)



03

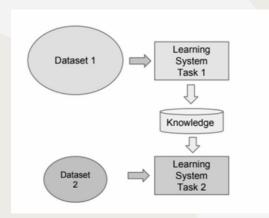
OUR APPROACH

PAST RESEARCH

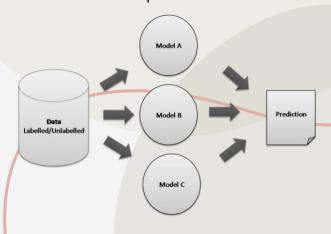
CNN — Transfer — Ensemble

Transfer learning models:

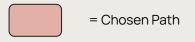
AlexNet, VGG, ResNet, Inception, Xception, ... The easier and faster aspect with pre-trained CNN model



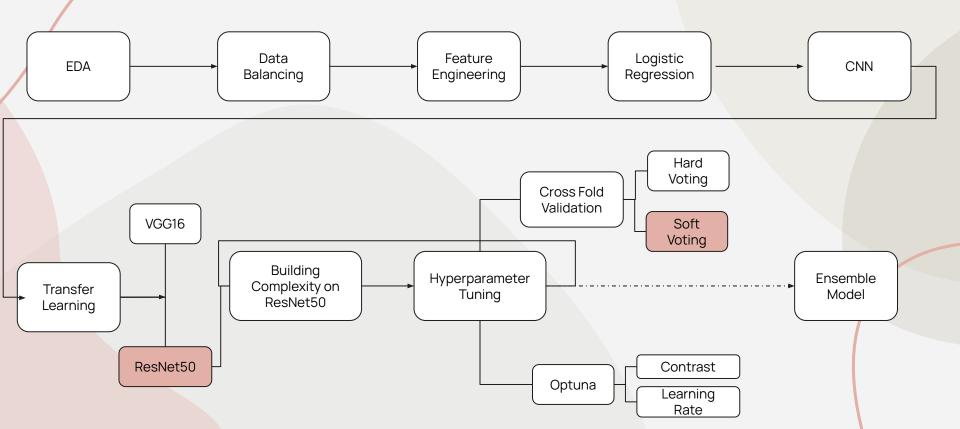
Reduce the generalization error of the prediction



APPROACH OVERVIEW



----- = Future Direction



EVALUATION METRICS

METRIC	RATIONALE	
Validation Accuracy	How well the model performs on validation data, validation accuracy is generally an indication if the model is overfitting or not.	
Test Accuracy	How well the model performs on test data.	
Validation False Negative Rate	How many true positives are wrongly misclassified as benign in the validation data	
Test False Negative Rate	How many true positives are wrongly misclassified as benign in the test data	
Validation AUC	FPR vs TPR rate, denotes how well the classifier can differentiate between benign and malignant class in the validation data, higher the number better it's	
Test AUC	FPR vs TPR rate, denotes how well the classifier can differentiate between benign and malignant class in the test data, higher the number better it's	

PERFORMANCE OF PREVIOUS MODEL EXPERIMENTS

Iterations Model		Number of Parameters	Test Accuracy	Test AUC
1	Logistic Regression	22,564,225	63.74%	64.51%
2	CNN	4,906,753	80.55%	80.56%
3	CNN with dropout layer	4,906,753	79.77%	79.28%
4	CNN with BatchNormalization	4,773,738	82.73%	80.75%
5	CNN, multiple convolutional layer and BatchNormalization with asymmetric filtering	4, 779, 530	83.74%	91.60%
6	VGG16	14, 714, 688	84.11%	84.28%
7	Mobilenet	3, 229, 889	91.15%	97.29%
8	Xception + Mobilenet	23, 122, 793	90.75%	97.19%

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FINAL MODEL

MODELS TESTED

VGG16



ResNet50

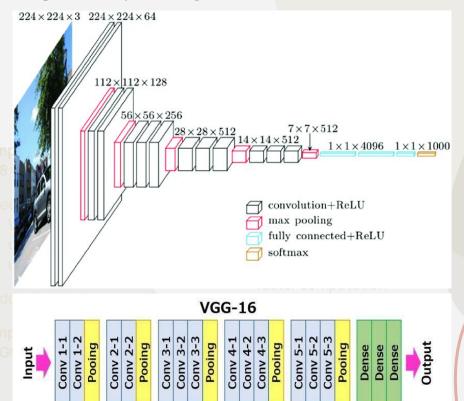


Xception & MobileNet

MODELS TESTED

VGG16

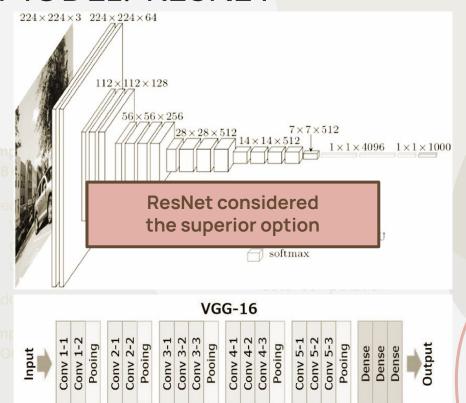
- 13 Conv2D + 3 Dense
- Small kernels (3x3 or 1x1):
 - multiple layers w/o pooling → mimics lager kernels
 - but more non-linearity & less parameters
- Increasing channel size
- Reduces effort on testing layer adding



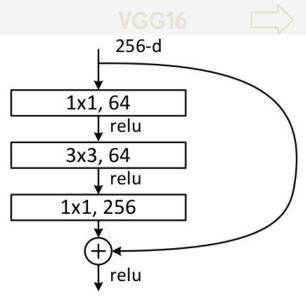
MODELS TESTED

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MODELS TESTED



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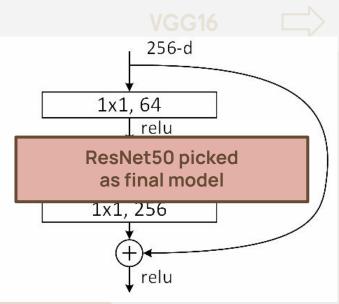


- Improved VGG, with
 48 Conv2D + 2 Pooling
- Deeper networks:
 - VGG shows deeper **better**
 - degradation can happen, unrelated to overfitting
- Adds shortcut connections
- Improved performance over VGG

- Improved Inception model
- main idea: multiple convolutions, then concatenating them
- Depthwise separable convolution layers for faster computation

Reference: Deep Residual Learning for Image Recognition; by K He et al; 2016

MODELS TESTED



ResNet50



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震 FINAL MODEL: ARCHITECTURE

BASE TRANSFER LEARNING MODEL

- ResNet50:

- 50 layers total
- 3 channels input
- at least 32 x 32
- trained on ImageNet

	·					
layer name	output size	18-layer	34-layer	50-layer	101-layer	152-layer
conv1	112×112			7×7, 64, stride 2		•
				3×3 max pool, strid	2	
conv2_x	56×56	$\left[\begin{array}{c} 3\times3,64\\ 3\times3,64 \end{array}\right]\times2$	$\left[\begin{array}{c} 3\times3, 64\\ 3\times3, 64 \end{array}\right]\times3$	$\begin{bmatrix} 1 \times 1, 64 \\ 3 \times 3, 64 \\ 1 \times 1, 256 \end{bmatrix} \times 3$	$\begin{bmatrix} 1 \times 1, 64 \\ 3 \times 3, 64 \\ 1 \times 1, 256 \end{bmatrix} \times 3$	$\begin{bmatrix} 1 \times 1, 64 \\ 3 \times 3, 64 \\ 1 \times 1, 256 \end{bmatrix} \times 3$
conv3_x	28×28	$\left[\begin{array}{c} 3\times3, 128\\ 3\times3, 128 \end{array}\right] \times 2$	$\left[\begin{array}{c} 3\times3, 128\\ 3\times3, 128 \end{array}\right]\times4$	$ \left[\begin{array}{c} 1 \times 1, 128 \\ 3 \times 3, 128 \\ 1 \times 1, 512 \end{array}\right] \times 4 $	$\begin{bmatrix} 1 \times 1, 128 \\ 3 \times 3, 128 \\ 1 \times 1, 512 \end{bmatrix} \times 4$	$ \left[\begin{array}{c} 1 \times 1, 128 \\ 3 \times 3, 128 \\ 1 \times 1, 512 \end{array}\right] \times 8 $
conv4_x	14×14	$ \begin{bmatrix} 3 \times 3, 256 \\ 3 \times 3, 256 \end{bmatrix} \times 2 $	$\left[\begin{array}{c} 3\times3,256\\ 3\times3,256 \end{array}\right]\times6$	$ \begin{bmatrix} 1 \times 1, 256 \\ 3 \times 3, 256 \\ 1 \times 1, 1024 \end{bmatrix} \times 6 $	$\begin{bmatrix} 1 \times 1, 256 \\ 3 \times 3, 256 \\ 1 \times 1, 1024 \end{bmatrix} \times 23$	$\begin{bmatrix} 1 \times 1, 256 \\ 3 \times 3, 256 \\ 1 \times 1, 1024 \end{bmatrix} \times 36$
conv5_x	7×7	$ \begin{bmatrix} 3 \times 3, 512 \\ 3 \times 3, 512 \end{bmatrix} \times 2 $	$\left[\begin{array}{c} 3\times3,512\\ 3\times3,512 \end{array}\right]\times3$	$ \begin{bmatrix} 1 \times 1, 512 \\ 3 \times 3, 512 \\ 1 \times 1, 2048 \end{bmatrix} \times 3 $	$ \begin{bmatrix} 1 \times 1, 512 \\ 3 \times 3, 512 \\ 1 \times 1, 2048 \end{bmatrix} \times 3 $	$ \left[\begin{array}{c} 1 \times 1, 512 \\ 3 \times 3, 512 \\ 1 \times 1, 2048 \end{array}\right] \times 3 $
	1×1 average pool, 1000-d tc, softmax					
FLO	OPs	1.8×10^9	3.6×10^9	3.8×10 ⁹	7.6×10^9	11.3×10^9

Source: Deep Residual Learning for Image Recognition; by K He et al; 2016

無 FINAL MODEL: ARCHITECTURE

FOR EACH K-FOLD:

- ResNet50

- After the 50 layers:

Global Average Pooling

- **Dropout** (rate at 0.1)

- Flatten

- Dense FC (2048 → 1; logits for soft voting)

All layers fully trainable, given:

Large dataset

Difference between medical & general images ===

- Optimizer: Adam with eta = 0.001

of epochs: 10

Model: "model"

Layer (type)	Output Shape	Param #
input_2 (InputLayer)	[(None, 96, 96, 3)]	0
resnet50 (Functional)	(None, 3, 3, 2048)	23587712
<pre>global_average_pooling2d (G lobalAveragePooling2D)</pre>	(None, 2048)	ø
dropout (Dropout)	(None, 2048)	0
flatten (Flatten)	(None, 2048)	0
dense (Dense)	(None, 1)	2049

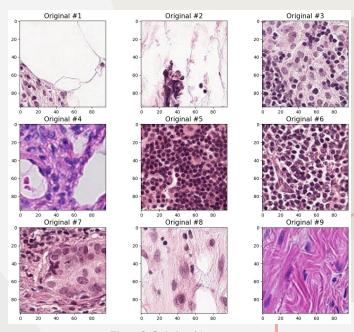
Total params: 23,589,761 Trainable params: 23,536,641 Non-trainable params: 53,120



FINAL MODEL: IMAGE PREPROCESSING

Image Augmentation Steps

- Unlike general image processing tasks, **medical images** (specifically histopathological tissue films) are:
 - more **standardised** (color, contrast, etc)
 - color carries no info (reflects dye color)
 - other random effects less likely (e.g. size changes, non-right-angled shifts, random shear, etc.)
 - Selected augmentations:
 - random flipping left-right
 - right-angled rotations (90°, 180° & 270°)
 - rotated images added to training set:
 - equally valid images
 - allows model to focus on patterns not depending on orientation



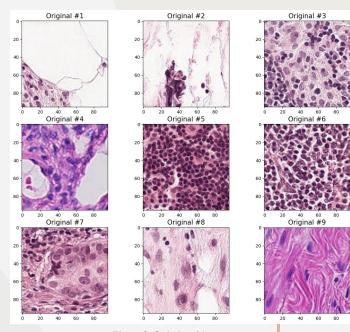
First 9 Original Images



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- Selected augmentations:
 - **grey-scaling** (then tripled it back to three channels)
 - random flipping & right-angled rotations (90°, 180° & 270°)
 - rotated images added to training set:
 - equally valid images
 - allows model to focus on patterns not depending on orientation



First 9 Original Images



FINAL MODEL: IMAGE PREPROCESSING

Image Augmentation Steps

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 - right-angled rotations (90°, 180° & 270°)
 - rotated images added to training set:
 - equally valid images
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- Consistent with experimentation

	Contrast Factor Applied to Images			
	0	3	5	
Avg Val Loss (Epoch 10)	0.3010 (sd: 0.046)	0.3908 (sd: 0.096)	0.3890 (sd: 0.144)	
Avg Val Acc (Epoch 10)	90.76% (sd: 1.43%)	89.28% (sd: 1.80%)	88.78% (sd: 3.63%)	
Test Acc (Ensemble)	94.48%	93.42%	92.32%	
AUC ROC (Ensemble)	0.9854	0.9805	0.9777	

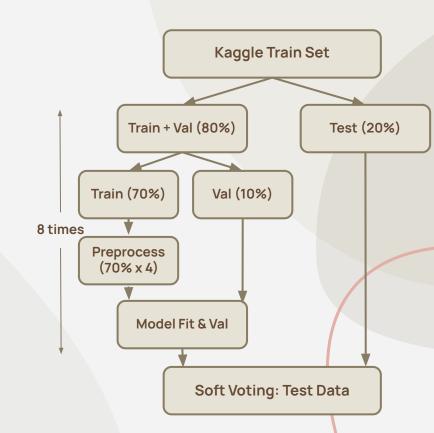
FINAL MODEL: K-FOLD & VOTING

K-FOLD PROCESS

- Stratified K-fold at k = 8:
 - K-value: tidy train-val-test split (7:1:2)
 - Stratified: balanced classes in train & test sets
 - Image preprocessing: on training set only

VOTING

- Soft-voting:
 - All 8-folds predict on test data
 - Final decision based on average probabilities
 - Consistently **better than hard voting**
 - lack of extreme outliers (models all similar)
 - Improvement of ~0.495 in AUC ROC score
 - Ensemble improves acc by ~3.69%, AUC by ~0.19°



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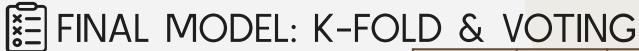
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	Voting Mechanism		Improvement:		
	Hard	Hard Soft			
Models with Conf	trast = 0				
- Test Acc	94.48%	94.52%	+0.04%		
- AUC ROC	0.9805	0.9854	+0.0049		
Models with Conf	Models with Contrast = 3				
- Test Acc	93.42%	93.47%	+0.05%		
- AUC ROC	0.9758	0.9805	+0.0048		
Models with Conf	Models with Contrast = 5				
- Test Acc	93.32%	92.60%	+0.29%		
- AUC ROC	0.9699	0.9777	+0.0079		

^{*} True positive and true negatives show strict improvements as well



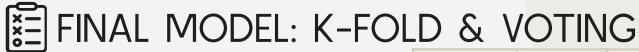
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- Ensemble improves acc by ~3.69%, AUC by ~1.91% (with diminishing returns)

	Single Model (Avg)	Ensemble Model	Improvement		
Models with Contrast = 0 (8 folds)					
- Test Acc	90.83% (0.013%)	94.52%	+3.69%		
- AUC ROC	0.9664 (0.009)	0.9854	+0.019		
Models with Contrast = 3 (8 folds)					
- Test Acc	89.47% (O.O13%)	93.47%	+4.00%		
- AUC ROC	0.9565 (0.009)	0.9805	+0.024		
Models with Contrast = 5 (8 folds)					
- Test Acc	89.02% (0.032%)	93.47%	+4.45%		
- AUC ROC	0.9523 (0.021)	0.9777	+0.025		

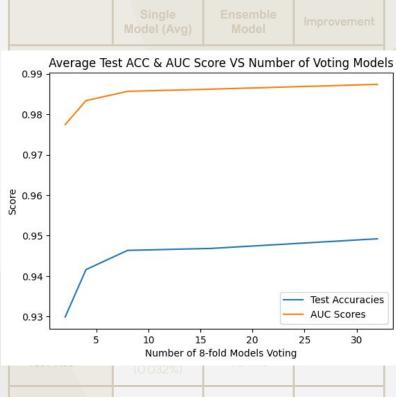


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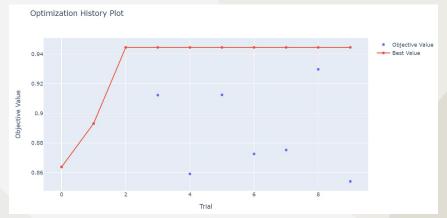


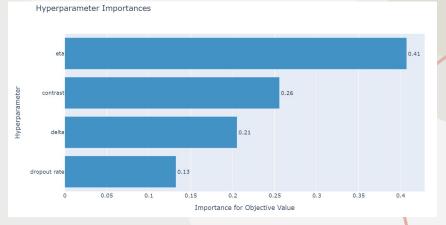
مهم

FINAL MODEL: OPTUNA

Model Tuning Using Optuna

- Optuna: using Bayesian model for hyperparameter tuning
- Study run with 10 trials, optimize on AUC:
 - Learning rate: 0.1 0.001, log-scale
 - **Dropout** rate: 0.1 0.5, float
 - Contrast factor: -1 5, integer
 - **Delta**: 0 0.5, float
 - Reduced image augmentation to 2x (from 4x) due to memory issues
- Best trial parameters:
 - Learning rate: 0.00121
 - **Dropout** rate: 0.2497
 - Contrast factor: -1
 - **Delta**: 0.2150
- More testing required for convergence





Learning Rates

- 4 sets of numbers tested:
 - 0.001 (Adam default)
 - 0.00121 (Optuna rate)
 - 0.01
 - 0.1
- At 10 epochs:
 - Best result: 0.001
 - Minor diff between 0.001 & 0.00121
 - More optuna trials may be needed
- Other issues:
 - Insufficient time to tune epoch num
 - Rate decay unexplored

	Learning Rate				
	0.001	0.00121	0.01	0.1	
Avg Val Loss (Epoch 10)	0.3010 (sd: 0.046)	O.2882 (sd: O.063)	0.3060 (sd: 0.042)	O.4134 (sd: O.136)	
Avg Val Acc (Epoch 10)	90.76% (sd: 1.43%)	90.93% (sd: 1.31%)	88.42% (sd: 2.03%)	83.03% (sd: 5.43%)	
Test Acc (Ensemble)	94.48%	94.16%	93.17%	90.08%	
AUC ROC (Ensemble)	0.9854	0.9830	0.9793	0.9777	

S FINAL MODEL: LEARNING RATE

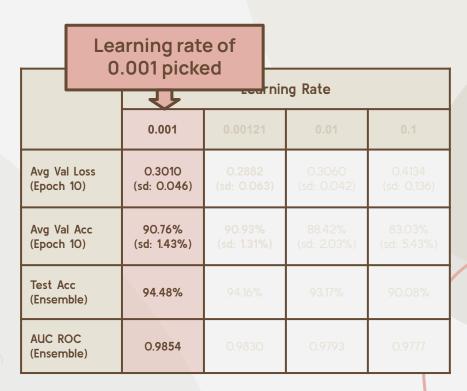
Learning Rates

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 - 0.001 (Adam default)
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 - 0.01
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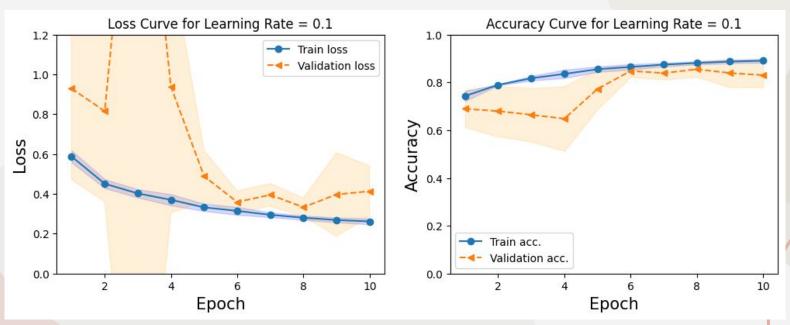
Rate decay unexplored





LEARNING RATE: 0.1

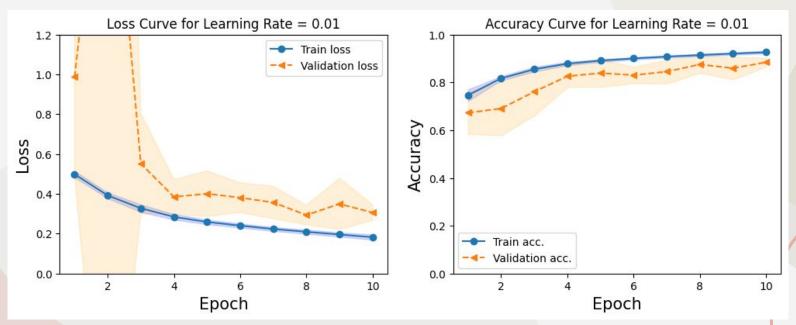
Loss and Accuracy Curves





LEARNING RATE: 0.01

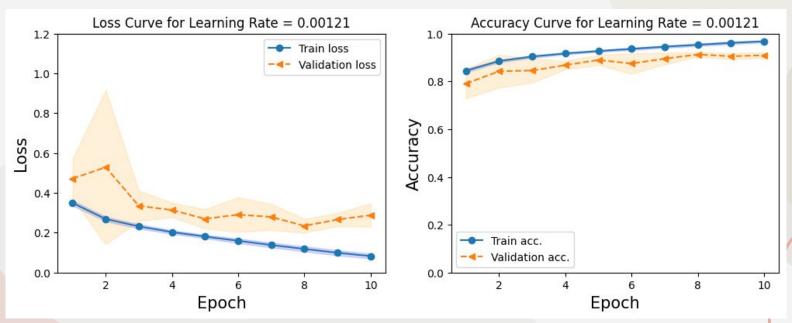
Loss and Accuracy Curves





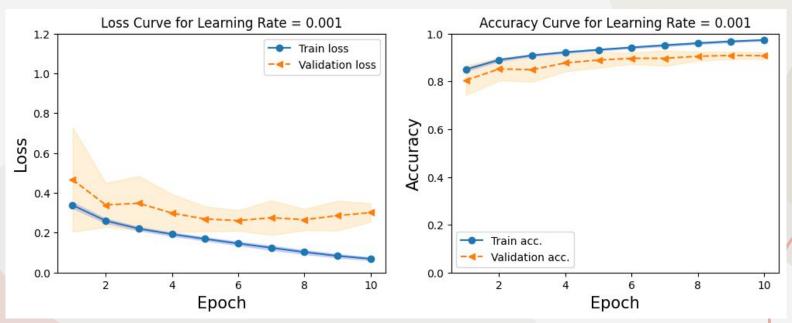
LEARNING RATE: 0.00121

Loss and Accuracy Curves





Loss and Accuracy Curves



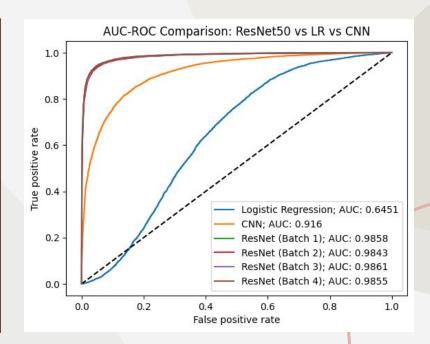
LEARNING RATES: CONFUSION MATRIX COMPARISON

	Consistently Goo		Learning Rate		
	0.001	0.00121	0.01	0.1	
True Positives	8307 (93%)	8273 (93%)	8337 (94%)	7452 (84%)	
False Negatives	604 (7%)	638 (7%)	574 (6%)	1459 (16%)	
True Negatives	8538 (96%)	85O9 (95%)	8267 (93%)	86O2 (97%)	
False Positives	373 (4%)	402 (5%)	644 (7%)	309 (3%)	



FINAL MODEL: PERFORMANCE SUMMARY

	Performance (over 4 batches)		
Overall test accuracy	94.48% (sd: 0.21%)		
- Precision	93.22% (sd: 0.47%)		
- Recall	95.71% (sd: 0.87%)		
F1 score	0.9445 (sd: 0.003)		
AUC-ROC score	0.9854 (sd: 0.0007)		



05

FUTURE WORK

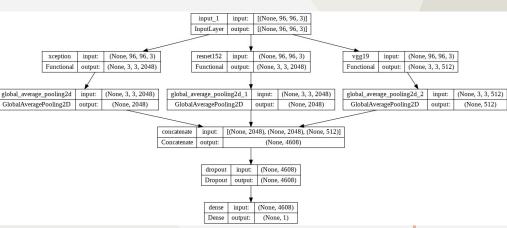


ENSEMBLE MODELS

Xception + MobileNet

[(None, 96, 96, 3)] input 1 [(None, 96, 96, 3)] InputLayer (None, 96, 96, 3) mobilenetv2 1.00 96 (None, 96, 96, 3) input: input: Functional output: (None, 3, 3, 2048) Functional output: (None, 3, 3, 1280) global_average_pooling2d (None, 3, 3, 2048) global_average_pooling2d_1 (None, 3, 3, 1280) input: GlobalAveragePooling2D (None, 2048) GlobalAveragePooling2D output: (None, 1280) input: [(None, 2048), (None, 1280)] concatenate Concatenate output: (None, 3328) (None, 3328) (None, 3328) (None, 3328) input: (None, 1)

Xception+ResNet+VGG19



06

REFERENCES

REFERENCES

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CONTRIBUTIONS

Chenyu Wang

- Codes & slides
- Literature review:
 - model
- architectures
- Data preprocessing
- Baseline model
- CNN model tuning
- PCA try and error
- Ensemble model:
 - Efficientnetb7 + inception+vgg19
- Brainstorming

Hector Rincon

Base Resnet model code Densenet201 exploration experiments Data preprocessing Code

deduplication/DRYing Slides

Optuna

experimentation

Ifrah Javed

- -Slides
- -Literature Review
- Ensemble Models
- Project Need
- -Baseline Model
- Optimization for
- LR Model
- GridSearch Optimization
- Learning Rate Optimization
- Brainstorming, general meeting planning, notes and setting up timeline and setting up team tools (Git, Google Folder, Presentation Slides)

- RESNET model:

Justin To

- tuning
- KPIs

- Experiments:

- K-fold & voting
- image preprocessing
- learning rates
- Optuna
- Preprocessing:
 - code review
- Literature review:
 - model architectures
- Brainstorming
- Codes & slides

Srila Maiti

Set up the initial structure of the code, reusable functions, classes, EDA, automating download of data from Kaggle

Baseline Logistic Regression Model, 5 CNN version experiments

Initial transfer learning model experiment with VGG16

Transfer Learning model experiments with Mobilnet, Restnet152, Xception and Mobilenet combination, InceptionRestnet, Densenet XGboost and image augmentation experiments and ensemble methods

Literature Review, coding, slides creation, brainstorming ideas

QUESTIONS?

THANK YOU!

07

BACKUP SLIDES

FINAL MODEL: OVERVIEW



Model Architecture

- ResNet50
- Trainable parameters: ~23.54 million



lmage Preprocessing

- Greyscale, then duplicate by 3 (ResNet takes in 3 channels)
- Random flipping
- All images rotated and added to original (i.e. 4x training size)



K-fold & Voting

- 8-folds, i.e. train-validation-test split is 7-1-2
- Train data then multiplied by 4 (i.e. total = $62,380 \times 4 = \sim 249,520$)
 - Soft-voting (i.e. average probabilities, not average votes)

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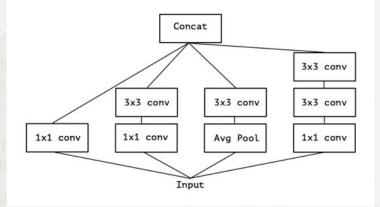
FINAL MODEL: RESNET

MODELS TESTED

- 13 Conv2D

Small kerne
- multiple la
pooling -kernels
- but more
less para

Figure 1. A canonical Inception module (Inception V3).



Xception & MobileNet

- Improved Inception model:
 - main idea: multiple convolutions, then concatenating them
- Depthwise separable convolution layers for faster computation

Reference: Xception: deep learning with depthwise separable convolutions; by F Chollet; 2016

FINAL MODEL: RESNET

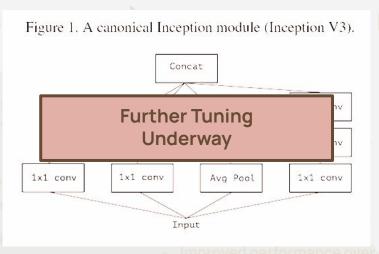
MODELS TESTED

- 13 Conv2D

- Small kernel multiple la pooling → kernels

- but more

Increasing



Xception

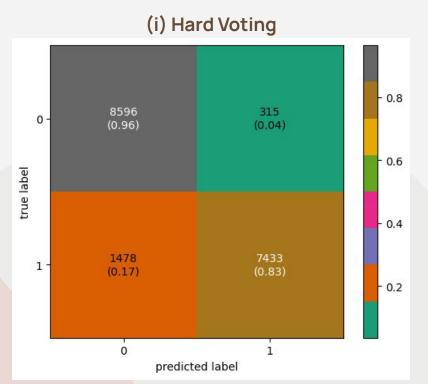
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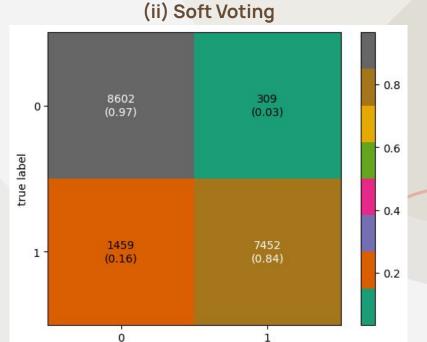
Reference: Xception: deep learning with depthwise separable convolutions; by F Chollet; 2016

layer adding



Confusion Matrix

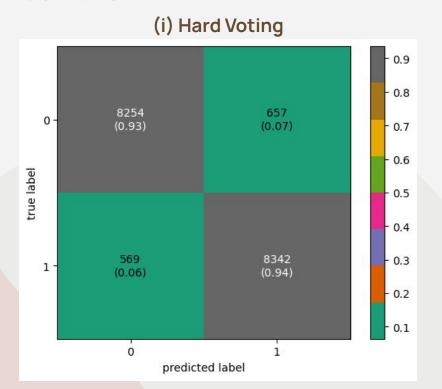


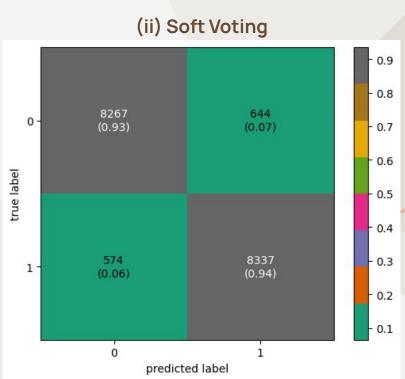


predicted label



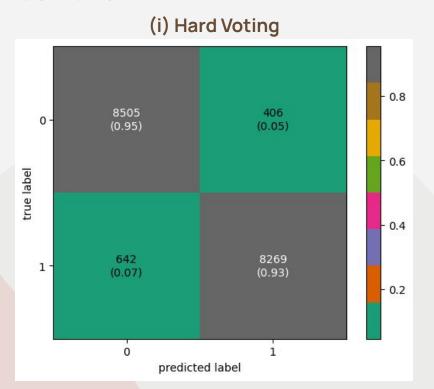
Confusion Matrix

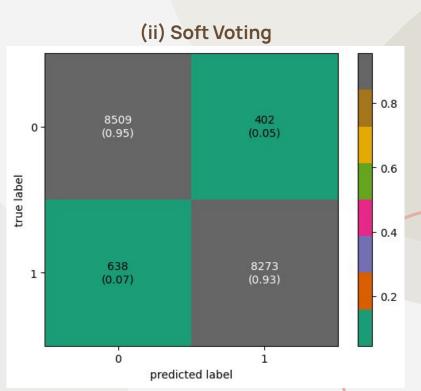






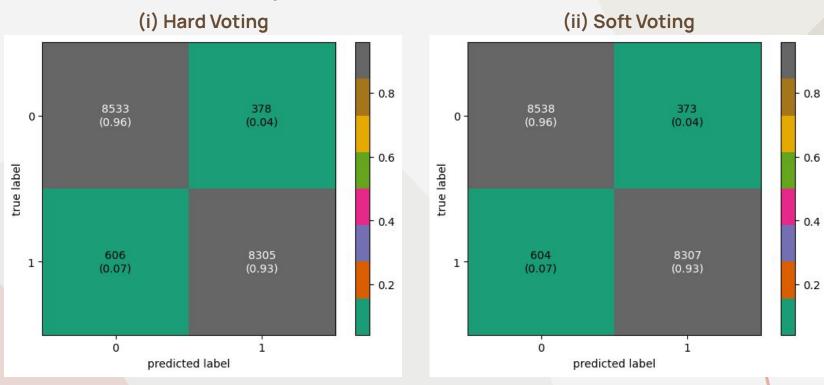
Confusion Matrix







Confusion Matrix (Average of 4 batches of 8 models)



DIFFERENT TRANSFER LEARNING MODELS EVALUATION

Available models							
Model	Size (MB)	Ton 1 Accumo	Top-5 Accura	Davametera	Depth	Time (me) no	Time (ms) per inference step (GPU)
Xception	Size (MB)	79.00%	94.50%		Depth 81	109.4	
VGG16	528	79.00%		138.4M	16	69.5	8.1
			\$2000 CO. CO.				4.2
VGG19	549	71.30%	27,070,040,000,175	143.7M	19	84.8	4.4
ResNet50	98	74.90%	92.10%		107	58.2	4.6
ResNet50V2	98	76.00%	93.00%		103	45.6	4.4
ResNet101	171	76.40%	92.80%		209	89.6	5.2
ResNet101V2	171	77.20%	93.80%		205	72.7	5.4
ResNet152	232	76.60%	93.10%		311	127.4	6.5
ResNet152V2	232	78.00%	94.20%		307	107.5	6.6
InceptionV3	92	77.90%	93.70%	7,001,001,001,000,001	189	42.2	6.9
InceptionResNetV2	215	80.30%	95.30%	747.472-2772-77	449	130.2	10
<u>MobileNet</u>	16	70.40%	89.50%	4.3M	55	22.6	3.4
MobileNetV2	14	71.30%	90.10%	3.5M	105	25.9	3.8
DenseNet121	33	75.00%	92.30%	8.1M	242	77.1	5.4
DenseNet169	57	76.20%	93.20%	14.3M	338	96.4	6.3
DenseNet201	80	77.30%	93.60%	20.2M	402	127.2	6.7
NASNetMobile	23	74.40%	91.90%	5.3M	389	27	6.7
NASNetLarge	343	82.50%	96.00%	88.9M	533	344.5	20
EfficientNetB0	29	77.10%	93.30%	5.3M	132	46	4.9
EfficientNetB1	31	79.10%	94.40%	7.9M	186	60.2	5.6
EfficientNetB2	36	80.10%	94.90%	9.2M	186	80.8	6.5
EfficientNetB3	48	81.60%	95.70%	12.3M	210	140	8.8
EfficientNetB4	75	82.90%	96.40%	19.5M	258	308.3	15.1
EfficientNetB5	118	83.60%	96.70%	30.6M	312	579.2	25.3
EfficientNetB6	166	84.00%	96.80%	43.3M	360	958.1	40.4
EfficientNetB7	256	84.30%	97.00%	66.7M	438	1578.9	61.6