



PRACTICUM II

Final Presentation

Deep Learning for Image Detection of Breast Cancer

Angela De La Mora
Regis University
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Project Overview

- Binary classification: cancer vs. no-cancer tissue in medical images
- Convolutional Neural Network architecture (aka: CovNet / CNN)
- Evolutionary algorithms to find optimal hyperparameters
- Total image count: 277,524
- Dataset based on seminal paper published in 2014
- Dataset downloaded from a republishing blog/author (Janowczyk, 2015)
- Comparison with original researchers Cruz-Roa et al. (2014):

	Precision	Recall	F-Score	Specificity	Balanced Accuracy
Orig. paper:	0.6540	0.7960	0.7180	0.8886	0.8423
Our project:	0.7617	0.7926	0.7768	0.9017	0.8471

Project Steps

- **W1** - Project research, selection and writing of proposal.
- **W2** - Prepare hardware/software on local PC.
- **W3** - Data gathering, cleaning and splitting (training, validation and test sets).
- **W4** - Exploratory architecture development; exploratory model training.
- **W5** - Model training; further iterations in architecture and hyperparameters.
- **W6** - Fine tune model training.
- **W7** - Compare performance metrics and finalize model selection.
- **W8** - Build presentation slides, upload to GitHub and present project.

Hardware and software requirements

Hardware:

- GPU: Nvidia 1080 Ti / 3584 cores / 11 GB RAM
- CPU: AMD Ryzen 2700X / 3.7 GHz / 8 cores / 16 threads
- RAM: 16 GB

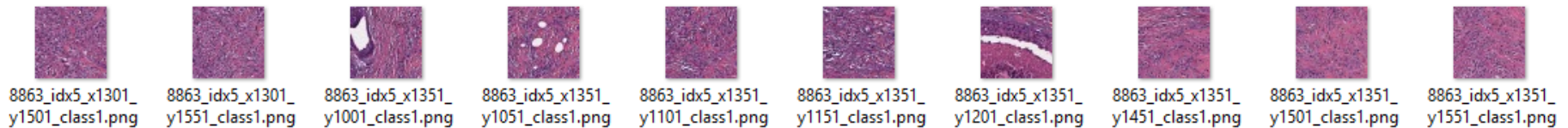
Software:

- Anaconda / Python 3.6
- Tensorflow 1.1 / Keras
- Nvidia Cuda 9.0 / cuDNN
- Jupyter Notebook

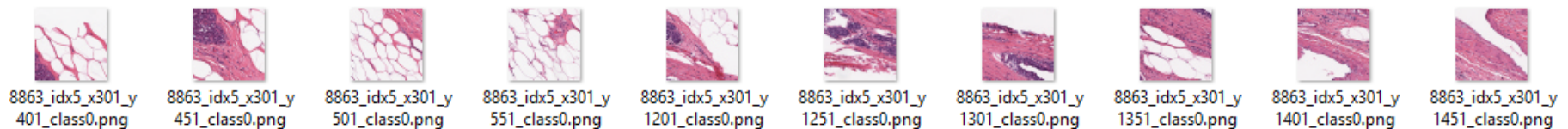


Data collection and preparation

- Image counts:
 - Negative class: 198,738
 - Positive class: 78,786
 - Total: 277,524
- Dataset size: 1.6 GB
- Image size: 50 x 50 pixels
- Mount slides of breast cancer specimens at 40x magnification
- Example positive class images (cancer):



- Example negative class images (no cancer):



Data collection and preparation

Categories:

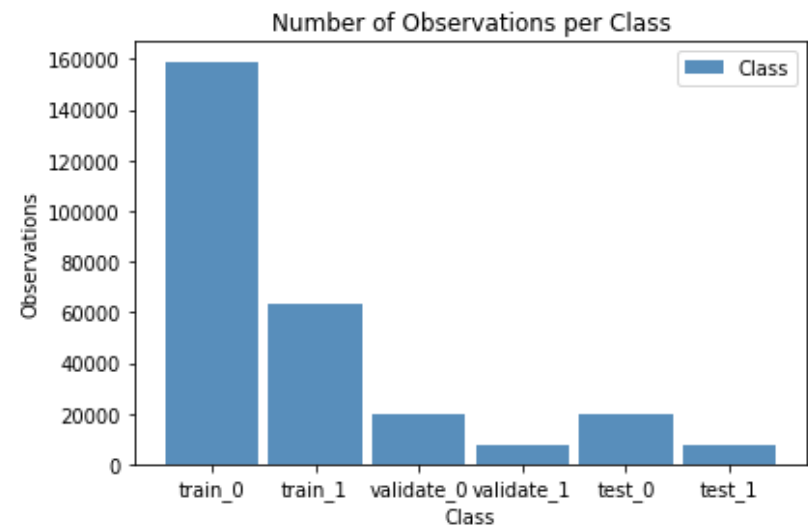
- Negative class: “0”
- Positive class: “1”

Data splitting by set, class and number of images:

- **Training set** – Class 0: 158,991 / Class 1: 63,029
- **Validation set** – Class 0: 19,874 / Class 1: 7,878
- **Test set** – Class 0: 19,873 / Class 1: 7,879

Highlights:

- Images selected at random
- Class imbalance present



Exploratory model architectures

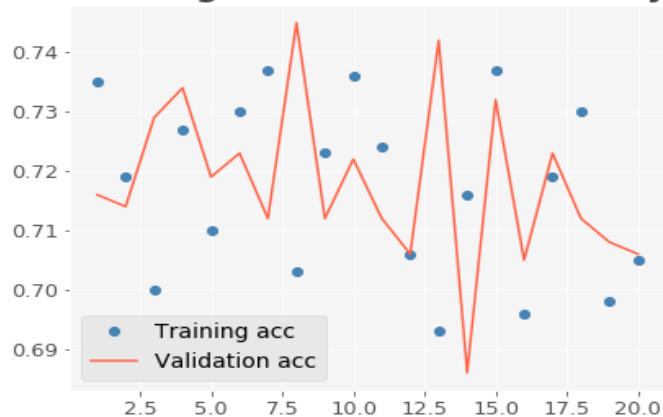
- Attempted a few simple model architectures to gauge performance
- Seven models compared
- **First model** not able to learn:

```
train_model(model_1, lr=0.01, epochs=20, epoch_steps=20, valid_steps=20, batch_size=50, image_size=(50, 50))
```

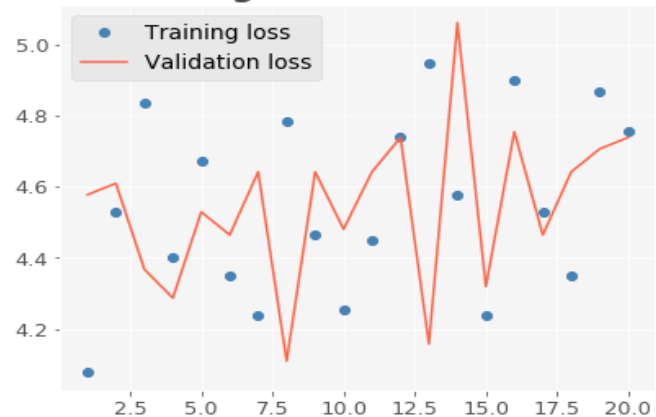
```
def model_1(lr, image_size):  
    model = Sequential()  
  
    model.add(Conv2D(32, (3, 3), activation='relu', input_shape=(image_size[0], image_size[1], 3)))  
  
    model.add(Conv2D(64, (3, 3), activation='relu'))  
  
    model.add(Flatten())  
  
    model.add(Dense(128, activation='relu'))  
    model.add(Dense(1, activation='sigmoid'))  
  
    model.compile(loss='binary_crossentropy', optimizer=Adam(lr=lr), metrics=['acc'])  
    model.summary()  
    return model
```

Layer (type)	Output Shape	Param #
conv2d_1 (Conv2D)	(None, 48, 48, 32)	896
conv2d_2 (Conv2D)	(None, 46, 46, 64)	18496
flatten_1 (Flatten)	(None, 135424)	0
dense_1 (Dense)	(None, 128)	17334400
dense_2 (Dense)	(None, 1)	129
Total params: 17,353,921		
Trainable params: 17,353,921		
Non-trainable params: 0		

Training and validation accuracy



Training and validation loss



Exploratory model architectures

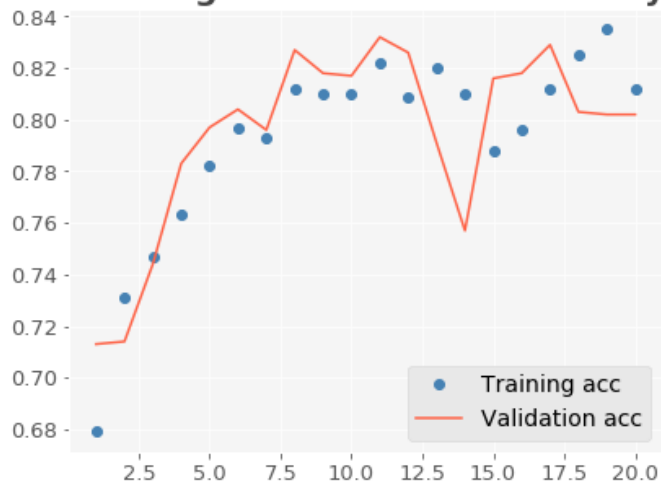
- Hyperparameter combinations were tweaked
- **The 3rd model** shows some improvement:

```
train_model(model_3, lr=0.0001, epochs=20, epoch_steps=20, valid_steps=20, batch_size=50, image_size=(50, 50))
```

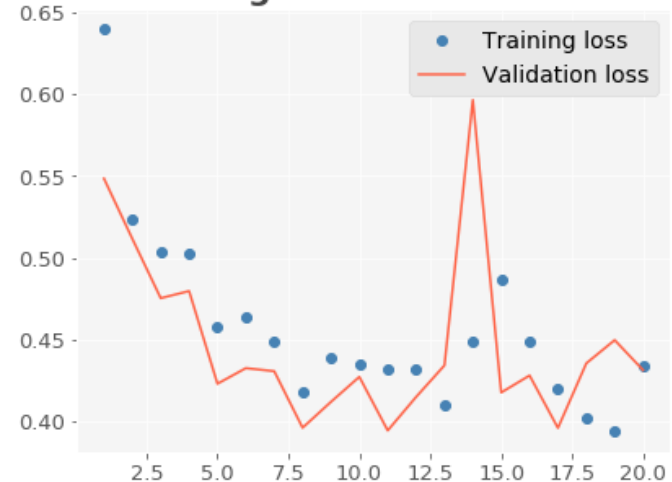
```
def model_3(lr, image_size):  
    model = Sequential()  
  
    model.add(Conv2D(32, (3, 3), activation='relu', input_shape=(image_size[0], image_size[1], 3)))  
  
    model.add(Conv2D(64, (3, 3), activation='relu'))  
  
    model.add(Flatten())  
  
    model.add(Dense(128, activation='relu'))  
    model.add(Dense(1, activation='sigmoid'))  
  
    model.compile(loss='binary_crossentropy', optimizer=Adam(lr=lr), metrics=['acc'])  
    model.summary()  
    return model
```

Layer (type)	Output Shape	Param #
conv2d_5 (Conv2D)	(None, 48, 48, 32)	896
conv2d_6 (Conv2D)	(None, 46, 46, 64)	18496
flatten_3 (Flatten)	(None, 135424)	0
dense_5 (Dense)	(None, 128)	17334400
dense_6 (Dense)	(None, 1)	129
Total params: 17,353,921		
Trainable params: 17,353,921		
Non-trainable params: 0		

Training and validation accuracy



Training and validation loss

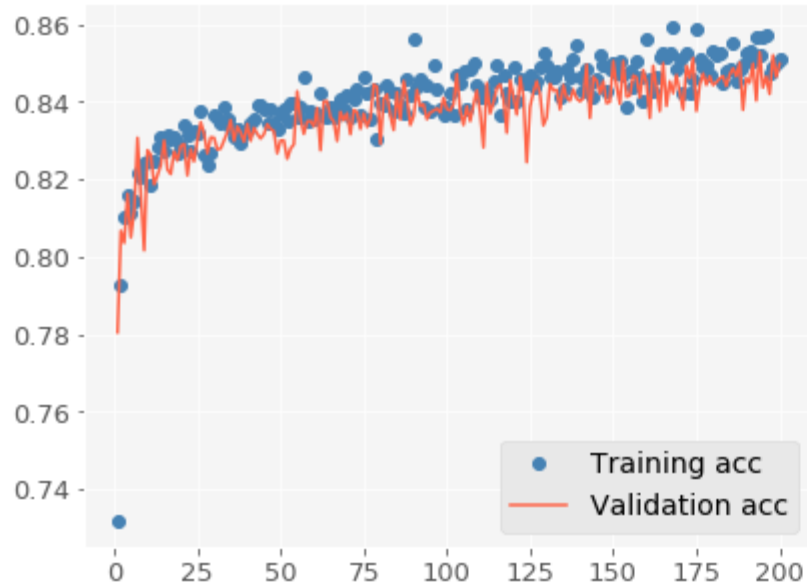


Exploratory model architectures

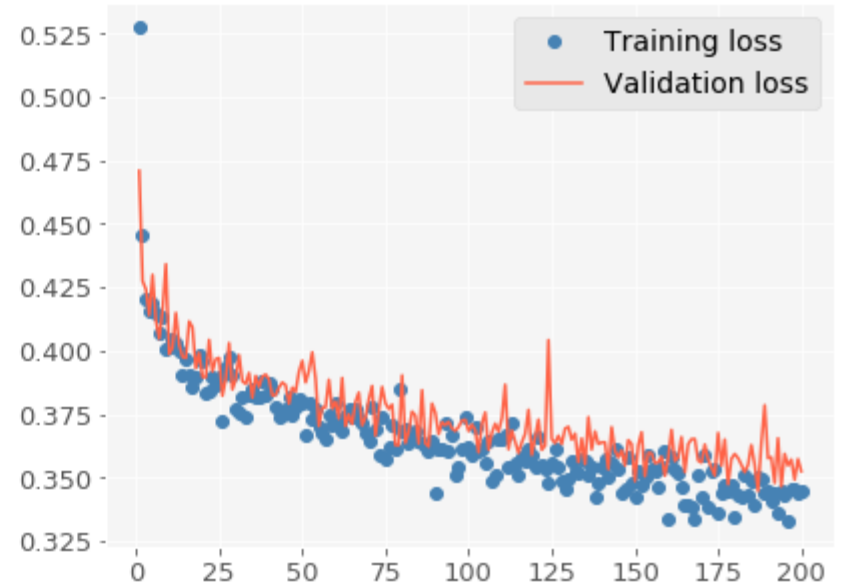
The 6th model starts showing signs of overfitting, where the training accuracy is consistently higher than the validation accuracy. The same applies to the training loss, which is consistently lower than validation loss:

```
train_model(model_6, lr=0.00001, epochs=200, epoch_steps=200, valid_steps=200, batch_size=50, image_size=(50, 50))
```

Training and validation accuracy



Training and validation loss



Exploratory model architectures

The 7th model, incorporates more layers and regularization (dropout) to combat overfitting by reducing the number of parameters from 17 million to about 300K.

```
train_model(model_7, lr=0.001, epochs=40, epoch_steps=140, valid_steps=10, batch_size=60, image_size=(50, 50))
```

```
def model_7(lr, image_size):
    model = Sequential()

    model.add(Conv2D(32, (3, 3), activation='relu', input_shape=(image_size[0], image_size[1], 3)))
    model.add(MaxPooling2D((2, 2)))

    model.add(Conv2D(64, (3, 3), activation='relu'))
    model.add(MaxPooling2D((2, 2)))

    model.add(Conv2D(128, (3, 3), activation='relu'))
    model.add(MaxPooling2D((2, 2)))

    model.add(Conv2D(128, (3, 3), activation='relu'))
    model.add(MaxPooling2D((2, 2)))

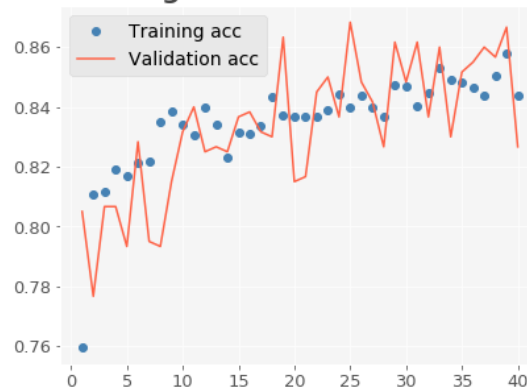
    model.add(Flatten())
    model.add(Dropout(0.5))

    model.add(Dense(512, activation='relu'))
    model.add(Dense(1, activation='sigmoid'))

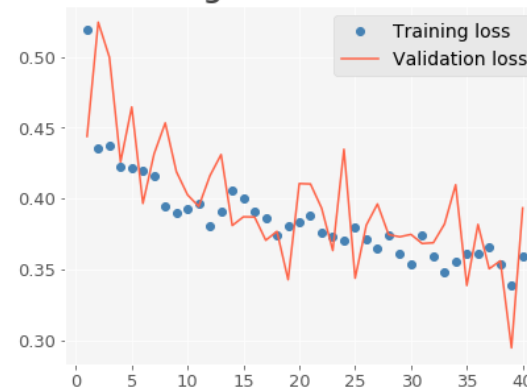
    model.compile(loss='binary_crossentropy', optimizer=Adam(lr=lr), metrics=['acc'])
    model.summary()
    return model
```

Layer (type)	Output Shape	Param #
conv2d_13 (Conv2D)	(None, 48, 48, 32)	896
max_pooling2d_1 (MaxPooling2D)	(None, 24, 24, 32)	0
conv2d_14 (Conv2D)	(None, 22, 22, 64)	18496
max_pooling2d_2 (MaxPooling2D)	(None, 11, 11, 64)	0
conv2d_15 (Conv2D)	(None, 9, 9, 128)	73856
max_pooling2d_3 (MaxPooling2D)	(None, 4, 4, 128)	0
conv2d_16 (Conv2D)	(None, 2, 2, 128)	147584
max_pooling2d_4 (MaxPooling2D)	(None, 1, 1, 128)	0
flatten_7 (Flatten)	(None, 128)	0
dropout_1 (Dropout)	(None, 128)	0
dense_13 (Dense)	(None, 512)	66048
dense_14 (Dense)	(None, 1)	513
Total params: 307,393		
Trainable params: 307,393		
Non-trainable params: 0		

Training and validation accuracy



Training and validation loss



Evolutionary algorithms

- A systematic approach to find an optimal neural network model requires exploring a large combination of architectures and hyperparameters, which is very difficult to do manually (Miikkulainen et al., 2019).
- Exploring 5 hyperparameter combinations (learning rate, number of epochs, steps per epoch, validation steps, and batch size) within known reasonable value bounds yields 100K possible models.
- Adding model architecture exploration (number of layers, nodes per layer, regularization, pooling, activation functions, etc.) yields a combinatorial explosion of over 10 billion models.
- Using 1,000 GPU's and a conservative average time of 5 minutes to train each model would require almost a million hours, or about 100 years to explore the whole solution space using brute force with current technology!
- Evolutionary algorithms use concepts from natural selection processes, especially neuroevolution, and applies them to the process of breeding the most fit models over the course of multiple generations until the best model is found for the task at hand (Stanley et al., 2019).

Evolutionary hyperparameter selection

- Use evolutionary algorithms instead of brute force or manual exploration

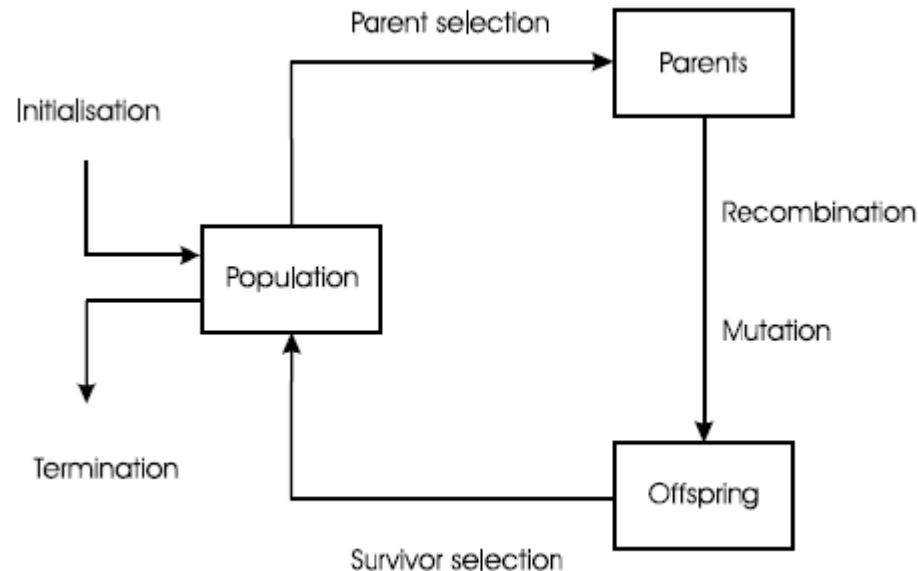


Fig. 1. Flowchart of evolutionary algorithm development (Eiben & Smith, 2015)

- Due to time constraints, only explore hyperparameters
- Use best model architecture found during model exploration (model 7)
- Hyperparameter values explored:
 - Learning rate: 5 values (0.1, 0.01, 0.001, 0.0001, 0.00001)
 - Batch size: 10 values (20, 40, 60, 80, 100, 120, 140, 160, 180, 200)
 - Epochs: 10 values (10, 20, 30, ..., 80, 90, 100)
 - Steps per epoch: 20 values (10, 20, 30, ..., 180, 190, 200)
 - Validation steps: 10 values (10, 20, 30, ..., 80, 90, 100)

Evolutionary hyperparameter selection

A **first generation** of 1,000 algorithms was initialized with values selected at random

```
# Create first generation  
first_generation(1000).tail()
```

	batch_size	steps_per_epoch	validation_steps	epochs	learning_rate	train_loss	train_acc	val_loss	val_acc	time	gen	alive
995	200	170	90	50	0.001						1	True
996	160	140	10	80	0.001						1	True
997	160	170	100	80	0.100						1	True
998	180	190	100	70	0.001						1	True
999	140	40	10	50	0.001						1	True

Survival of the fittest: After training, only the top 10% performing algorithms was selected

```
# Keep top performing algorithms  
df_fittest = select_fittest(top_percent=10)  
df_fittest.nlargest(10, 'val_acc')
```

	batch_size	steps_per_epoch	validation_steps	epochs	learning_rate	train_loss	train_acc	val_loss	val_acc	time	gen	alive
653	100	190	10	90	0.0001	0.317655	0.865789	0.292571	0.894000	0:11:41	1	True
994	140	180	50	50	0.0010	0.320372	0.864048	0.315853	0.874857	0:10:14	1	True
693	140	160	10	80	0.0010	0.315224	0.867411	0.319296	0.874613	0:13:23	1	True
605	60	200	30	80	0.0010	0.326009	0.861417	0.316879	0.873889	0:19:31	1	True
760	200	190	40	100	0.0001	0.298031	0.874026	0.304660	0.873125	0:39:18	1	True
963	140	200	90	90	0.0001	0.312412	0.867250	0.306135	0.872778	0:34:51	1	True
548	80	200	30	100	0.0010	0.313608	0.866063	0.324402	0.872083	0:21:46	1	True
996	160	140	10	80	0.0010	0.309834	0.872634	0.339324	0.871875	0:12:31	1	True
276	100	170	50	80	0.0001	0.335177	0.857294	0.311656	0.871600	0:24:30	1	True
215	180	130	90	100	0.0010	0.304454	0.868462	0.312375	0.871169	0:37:29	1	True

Evolutionary hyperparameter selection

- For the **second generation**, surviving algorithms were matched at random and their **chromosomes** (hyperparameters) passed to the next generation (**genetic inheritance**) with a 50% chance of inheriting a chromosome from either parent.
- Each set of parents produce 5 **offspring** at most, excluding offspring with chromosomal combinations previously observed in the population.
- A random chromosome **mutation** was assigned to one child per parent pair, promoting variety in the population and exploration of new values vs. exploitation of known best values.

```
# example of parents  
df_parents
```

	batch_size	steps_per_epoch	validation_steps	epochs	learning_rate	train_loss	train_acc	val_loss	val_acc	time	gen	alive
963	140	200	90	90	0.0001	0.312412	0.867250	0.306135	0.872778	0:34:51	1	True
923	80	110	50	90	0.0001	0.334619	0.858523	0.333084	0.854750	0:07:29	1	True

```
# example of children  
df_children
```

	batch_size	steps_per_epoch	validation_steps	epochs	learning_rate	train_loss	train_acc	val_loss	val_acc	time	gen	alive
0	80	200	90	90	0.0001						2	True
1	140	200	50	90	0.0001						2	True
2	80	110	50	90	0.0001						2	True
3	80	110	50	90	0.0001						2	True
4	140	200	90	90	0.1000						2	True

Evolutionary hyperparameter selection

- Best-performing algorithms from previous generations were allowed to survive if their validation accuracy fell within the top 10% best-performing algorithms in the population.
- A total of 4 generations were evolved
- A total of 17 algorithms survived the **mass extinctions**
- Total training time for all generations (1326 algorithms): almost 10 days (237.5 hours)
- Best-performing algorithms, based on validation accuracy:

	batch_size	steps_per_epoch	validation_steps	epochs	learning_rate	train_loss	train_acc	val_loss	val_acc	time	gen	alive
653	100	190	10	90	0.0001	0.317655	0.865789	0.292571	0.894000	0:11:41	1	True
1240	160	140	10	80	0.0010	0.309508	0.870714	0.298903	0.886250	0:12:35	3	True
1178	80	190	30	50	0.0010	0.333464	0.858750	0.292957	0.884583	0:05:13	2	True
1278	200	130	10	90	0.0010	0.302942	0.873192	0.296192	0.882000	0:17:44	3	True
1129	200	200	20	70	0.0001	0.311461	0.867375	0.295427	0.881832	0:29:50	2	True
1107	80	160	10	80	0.0010	0.321133	0.866250	0.336637	0.881250	0:10:42	2	True
1038	100	200	10	90	0.0010	0.306632	0.870800	0.317246	0.881000	0:14:31	2	True
1250	200	190	30	100	0.0010	0.284835	0.881026	0.298489	0.878500	0:28:42	3	True
1267	200	150	20	60	0.0010	0.310335	0.870633	0.298475	0.878250	0:13:40	3	True
1039	100	190	10	50	0.0001	0.344512	0.852526	0.317177	0.878000	0:07:40	2	True
1314	100	150	10	90	0.0010	0.311339	0.868067	0.298141	0.878000	0:12:12	4	True

- Some algorithms achieved comparable results in less time (i.e., #1178 vs. #1129)

Performance metrics

- Predictions on the test set images were run using all of the 17 surviving algorithms
- New performance metrics introduced based on predictions:
 - **Accuracy:** How many predictions were correct
 - **Precision:** How many positive predictions were true positives
 - **Sensitivity/Recall:** How many positives were correctly identified (true positive rate)
 - **Specificity:** How many negatives were true negatives
 - **F1-Score:** Weighted average of sensitivity and precision
- Accuracy may not be the best performance metric, especially when class imbalance exists
- Top performing models, based on F1-Score:

	model_name		cm	accuracy	precision	sensitivity	specificity	f1_score
0	1250	Predicted False True __all__\nActual ...		0.876766	0.790413	0.770148	0.919036	0.780149
1	1312	Predicted False True __all__\nActual ...		0.870712	0.761678	0.792613	0.901676	0.776838
2	1278	Predicted False True __all__\nActual ...		0.874423	0.791302	0.757457	0.920797	0.774009

- Top performing models, based on Sensitivity/Recall:

	model_name		cm	accuracy	precision	sensitivity	specificity	f1_score
0	1312	Predicted False True __all__\nActual ...		0.870712	0.761678	0.792613	0.901676	0.776838
1	1107	Predicted False True __all__\nActual ...		0.858605	0.736515	0.781571	0.889146	0.758374
2	653	Predicted False True __all__\nActual ...		0.864839	0.756908	0.771798	0.901726	0.764281

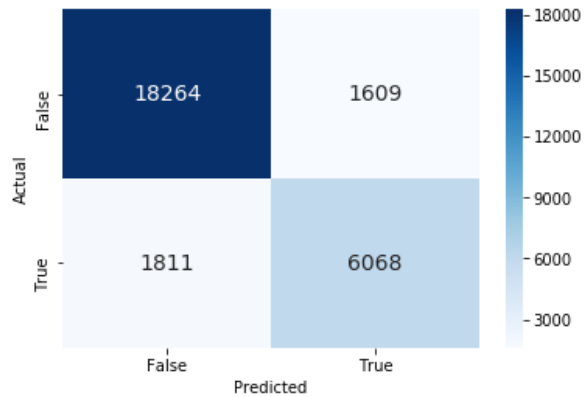
Performance metrics

Confusion Matrix Results



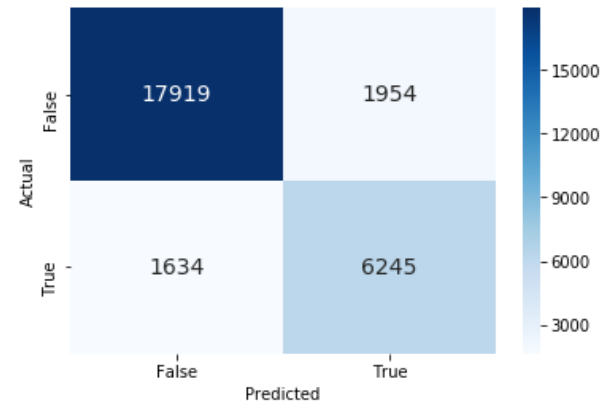
Best model based on F1-Score:

Predicted	False	True	__all__
Actual			
False	18264	1609	19873
True	1811	6068	7879
__all__	20075	7677	27752



Best model based on Sensitivity/Recall:

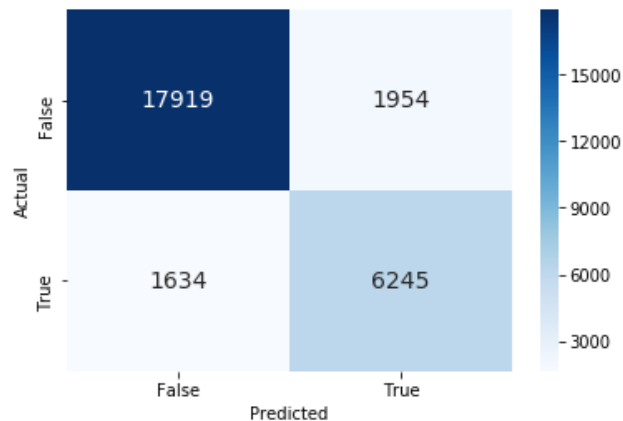
Predicted	False	True	__all__
Actual			
False	17919	1954	19873
True	1634	6245	7879
__all__	19553	8199	27752



Model selection

- Chosen model is the one based on highest sensitivity
- Maximizes true positives and minimizes false negatives
- Model based on F1-Score is a good balance, but our goal is to flag as many true positives as possible (here, false negatives are worse than false positives)

Example of predicted images →
(probabilities included for reference)



	file	y_true	y_pred	y_pred_prob
0	10253_idx5_x1001_y1501_class0.png	0	0	0.024449
1	10253_idx5_x1051_y151_class0.png	0	0	0.008627
2	10253_idx5_x1151_y551_class0.png	0	0	0.316427
3	10253_idx5_x1151_y701_class0.png	0	1	0.870326
4	10253_idx5_x1201_y1351_class0.png	0	0	0.187531
5	10253_idx5_x1201_y401_class0.png	0	0	0.041283
6	10253_idx5_x1201_y451_class0.png	0	1	0.656394
7	10253_idx5_x1251_y1251_class0.png	0	0	0.170695
8	10253_idx5_x1251_y901_class0.png	0	1	0.623490
9	10253_idx5_x1301_y1101_class0.png	0	1	0.760330

← **Confusion matrix of selected model**

Prediction results

True Negatives

17919

	file	y_true	y_pred	y_pred_prob
0	10253_idx5_x1001_y1501_class0.png	0	0	0.024449
1	10253_idx5_x1051_y151_class0.png	0	0	0.008627
2	10253_idx5_x1151_y551_class0.png	0	0	0.316427
3	10253_idx5_x1201_y1351_class0.png	0	0	0.187531
4	10253_idx5_x1201_y401_class0.png	0	0	0.041283

False Positives

1954

	file	y_true	y_pred	y_pred_prob
0	10253_idx5_x1151_y701_class0.png	0	1	0.870326
1	10253_idx5_x1201_y451_class0.png	0	1	0.656394
2	10253_idx5_x1251_y901_class0.png	0	1	0.623490
3	10253_idx5_x1301_y1101_class0.png	0	1	0.760330
4	10253_idx5_x1451_y1001_class0.png	0	1	0.759450

False Negatives

1634

	file	y_true	y_pred	y_pred_prob
0	10255_idx5_x151_y1051_class1.png	1	0	0.234125
1	10256_idx5_x1601_y1101_class1.png	1	0	0.217857
2	10256_idx5_x1851_y901_class1.png	1	0	0.443880
3	10256_idx5_x1951_y1401_class1.png	1	0	0.324580
4	10256_idx5_x2151_y951_class1.png	1	0	0.479504

True Positives

6245

	file	y_true	y_pred	y_pred_prob
0	10253_idx5_x601_y751_class1.png	1	1	0.649673
1	10253_idx5_x751_y451_class1.png	1	1	0.888613
2	10253_idx5_x751_y701_class1.png	1	1	0.706819
3	10253_idx5_x751_y751_class1.png	1	1	0.524811
4	10254_idx5_x1651_y1051_class1.png	1	1	0.906722

Prediction results

True Negatives



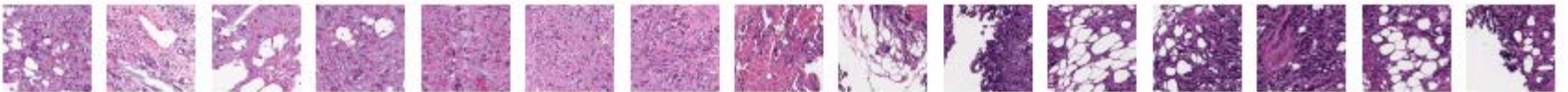
False Negatives



True Positives

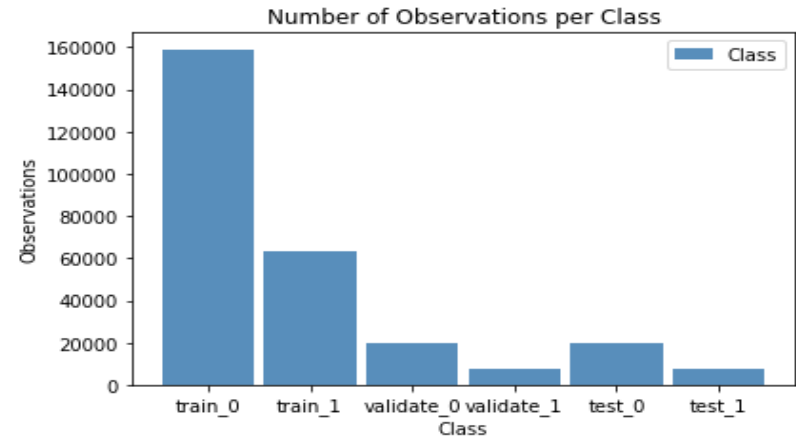


False Positives



Class imbalance and data augmentation

Class imbalance observed during data analysis



Weights can be added for class imbalance during training:

```
files_cnt = []
for category in categories:
    files = os.listdir(os.path.join(dir_train, category))
    files_cnt.append(int(len(files)))

class_weights = {}
for label in range(len(files_cnt)):
    class_weights.update({label : round(files_cnt[label]/sum(files_cnt)*100)})

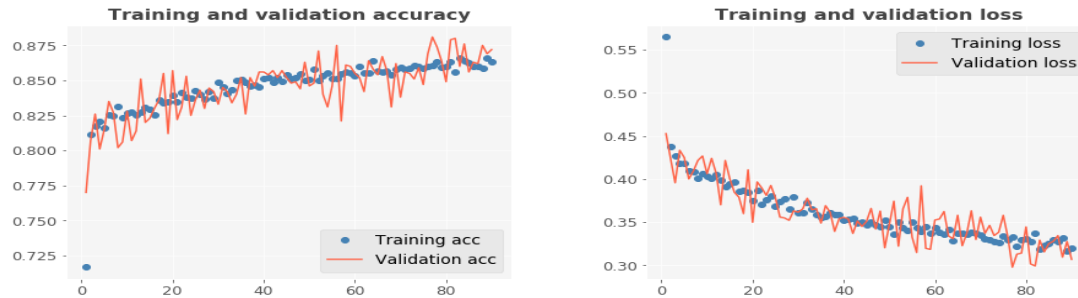
# start training
t0 = datetime.now()
with tf.device('/gpu:0'):
    history = model.fit_generator(gen_train, steps_per_epoch=epoch_steps, epochs=epochs, validation_data=gen_valid,
                                validation_steps=valid_steps, class_weight=class_weights)
```

Data augmentation can be added during training on the fly →

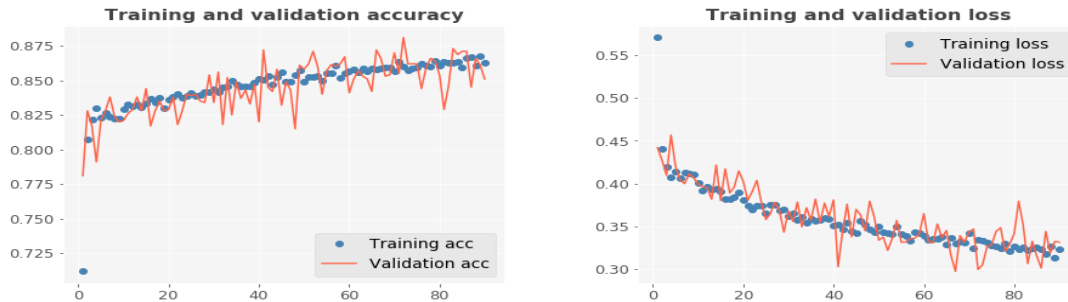
[illegible]

Class imbalance and data augmentation

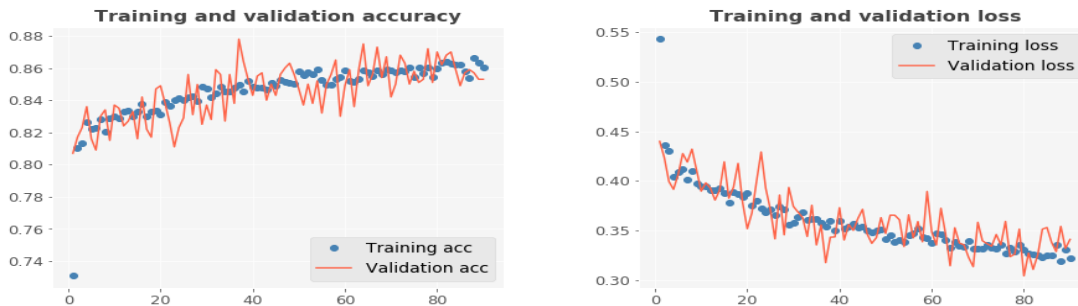
- Retrain best model with no imbalance correction or data augmentation (baseline model)



- Retrain best model with class imbalance correction only (no data augmentation)



- Retrain best model using class imbalance correction and data augmentation



Class imbalance and data augmentation

- Correcting for class imbalance did not yield to an improved model
- Applying data augmentation also had not positive effect
- The model without correction or augmentation performed better

model_name			cm	accuracy	precision	sensitivity	specificity	f1_score
0	2000	Predicted False True __all__\nActual ...		0.864586	0.771870	0.742480	0.912998	0.756890
1	2001	Predicted False True __all__\nActual ...		0.865884	0.779556	0.735626	0.917526	0.756954
2	2002	Predicted False True __all__\nActual ...		0.863649	0.792041	0.704785	0.926634	0.745870

Conclusions

- Evolutionary algorithms are a quick and effective alternative to brute force and manual training for deep neural networks.
- Binary class imbalance at ~ 70% / 30% did not negatively impact model performance, likely because the small class contains 63K images.
- Misclassified images show very close resemblance to the predicted class and error could be due to deficiencies in the original data labeling (this point was also raised on paper by original researchers).
- Performance metrics indicate similar results in sensitivity/recall, specificity and balanced accuracy to the baseline model by original researchers; and an important improvement in precision and F-score.

Future research

- Expand the use of evolutionary algorithms to include model architecture selection along with hyperparameters.
- To account for increased training time, extra GPU's and RAM could be added, or model could be trained on scalable cloud virtual machines.

References

- Cruz-Roa, A., et al. (2014). Automatic detection of invasive ductal carcinoma in whole slide images with Convolutional Neural Networks. Procedures of SPIE, 9041(904103). Retrieved from https://www.researchgate.net/publication/263052166_Automatic_detection_of_invasive_ductal_carcinoma_in_whole_slide_images_with_Convolutional_Neural_Networks
- Eiben, A. E., & Smith, J. E. (2015). Introduction to Evolutionary Computing, 2nd Ed. Amsterdam, The Netherlands: Springer. Print.
- Janowczyk, A. (2015). Invasive ductal carcinoma (IDC) segmentation. Retrieved from <http://www.andrewjanowczyk.com/use-case-6-invasive-ductal-carcinoma-idc-segmentation/>

Resources

Repository:

https://github.com/gelidely/CNN_Breast_Cancer_Detection

Software:

<https://www.tensorflow.org/>

<https://keras.io/>

<https://www.anaconda.com/>

<https://www.nvidia.com/en-us/geforce/products/10series/geforce-gtx-1080-ti/>

<https://developer.nvidia.com/cuda-downloads>

<https://developer.nvidia.com/cudnn>