

A Combined Radio-Histological Approach for Classification of Low Grade Gliomas

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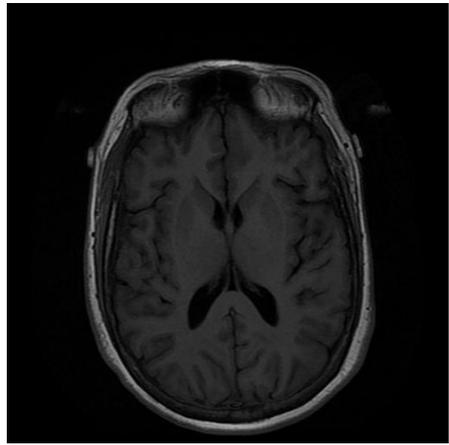
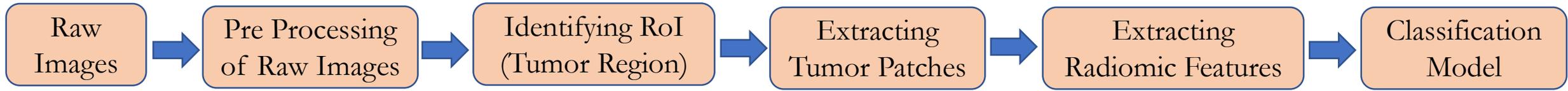


The logo for MiRL (Machine Intelligence Research Laboratory) consists of the letters 'MiRL' in a stylized font. The 'M' is blue, the 'i' is red, and the 'R' and 'L' are purple.

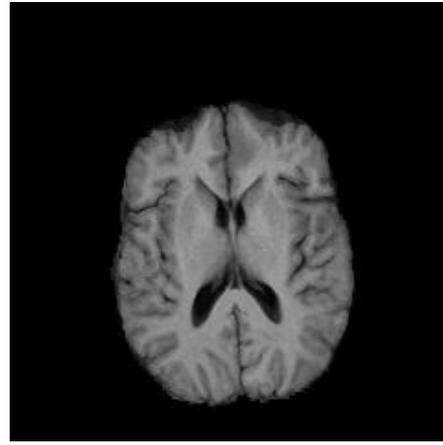
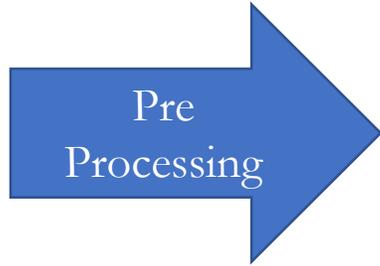
* Equal contribution

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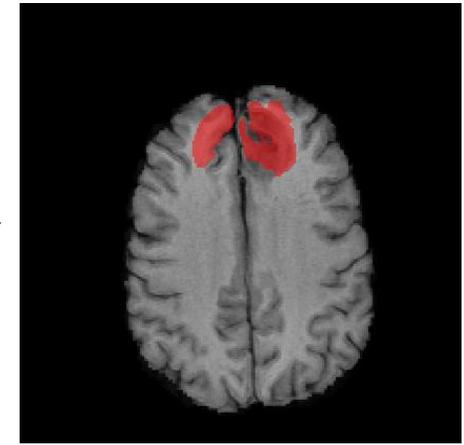
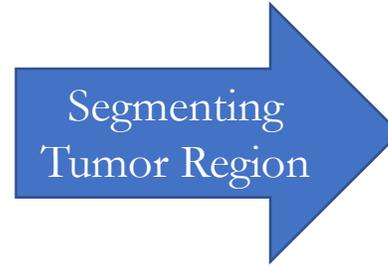
Radiology Pipeline



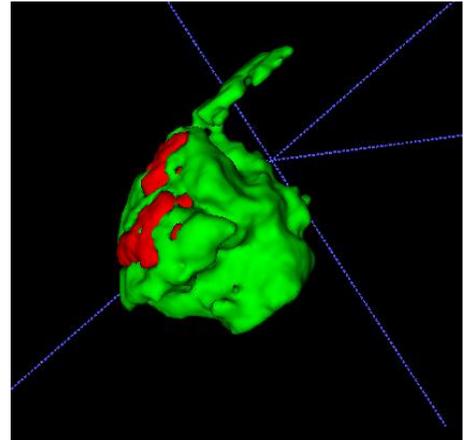
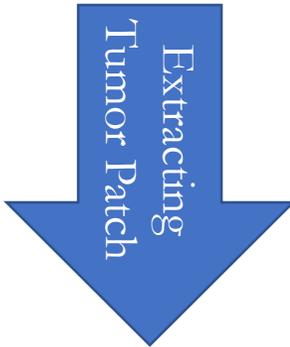
Raw MR Image



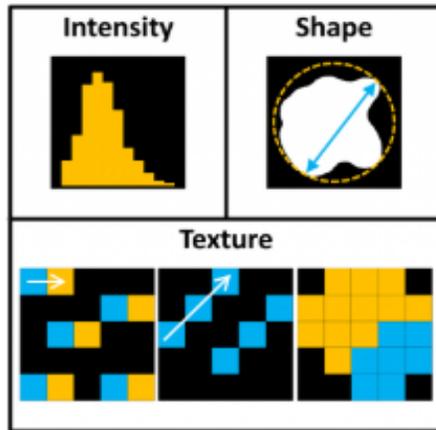
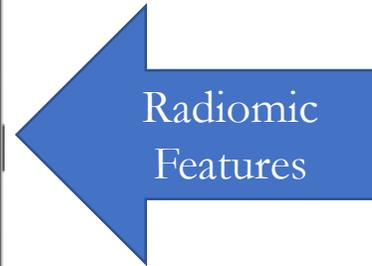
Pre Processed MR Image



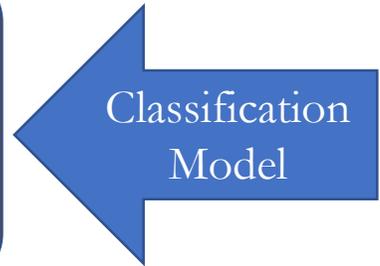
Tumor Segmentation



Tumor Patch Extraction



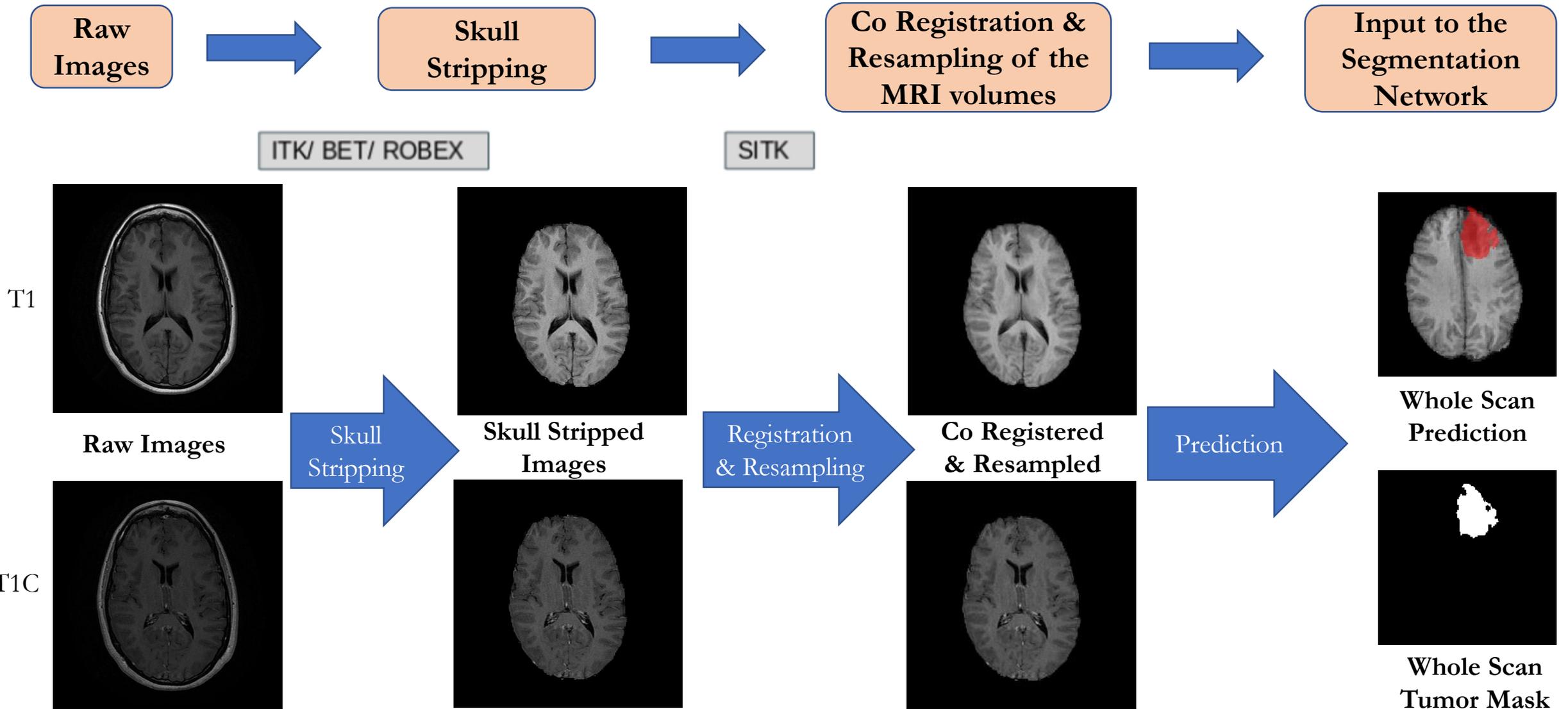
Radiomic Feature Extraction¹



$P(\text{Astrocytoma}) = ??$
 $P(\text{Oligodendroglioma}) = ??$

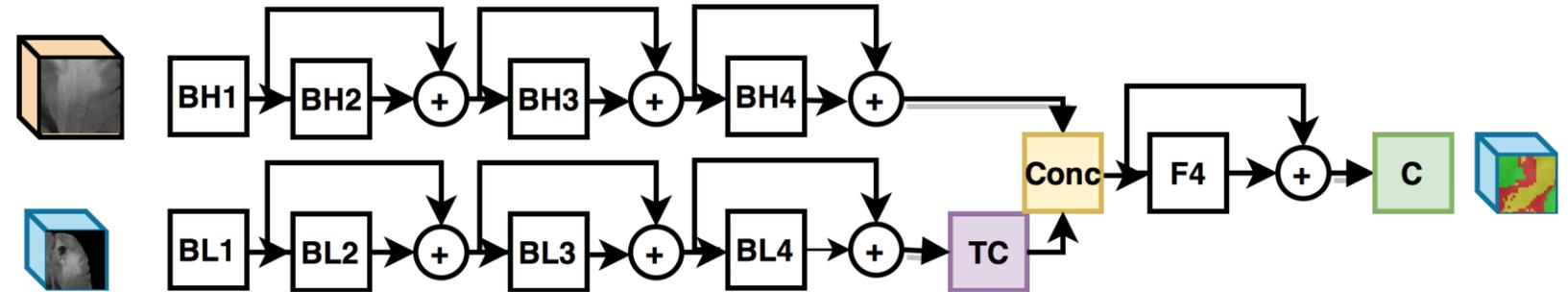
Pre Processing of Magnetic Resonance Images

The MR dataset were fed into a pre-processing pipeline which involved skull-stripping (ROBEX), co-registration of MR sequences to T1c and re-sampling of MR volumes to isotropic voxels.

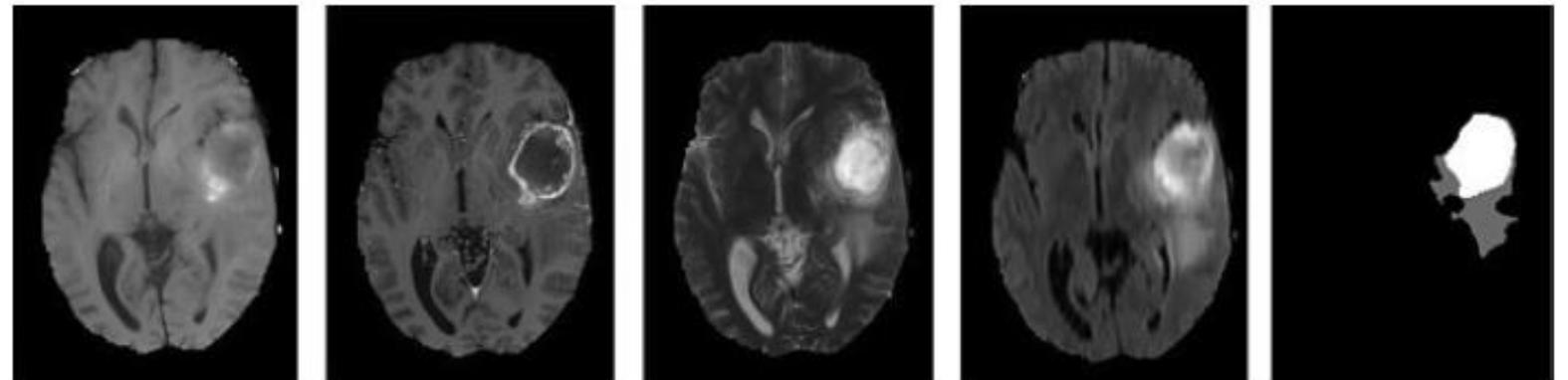


Segmentation of Brain Tumor

- A Fully Convolutional 3-D Neural Network was used to segment the tumor
- The network architecture extracts both local and global features from 3-D patches
- Network trained on publicly available MICCAI BraTS 2018 dataset
- Architecture inspired by Deep Medic¹ and Havaei et al²



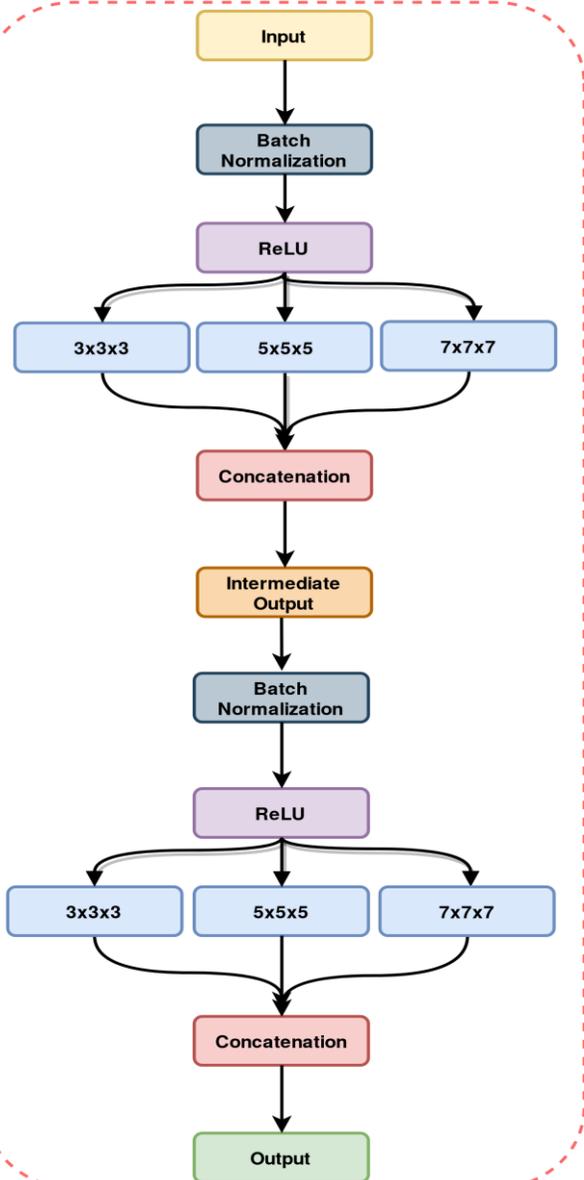
3-D Neural Network Architecture



From left to right: T1, T1C, T2, FLAIR, Segmented Tumor

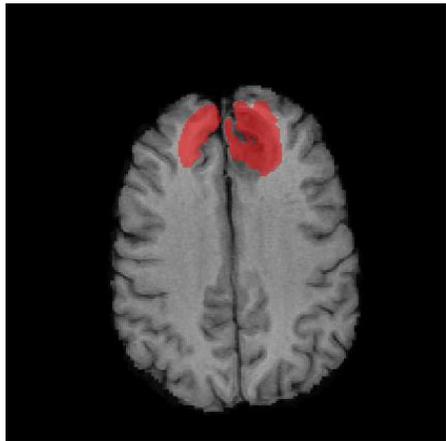
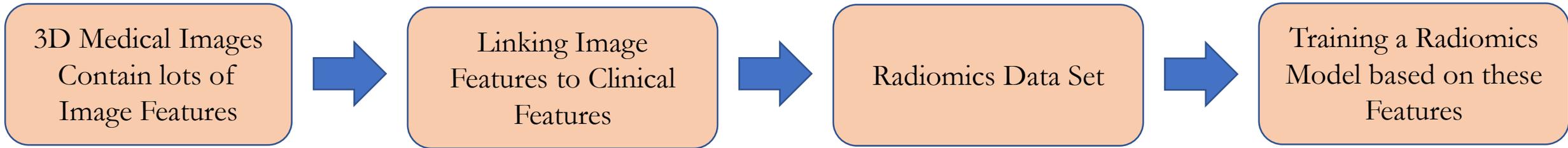
1. Kamnitsas K, Ledig C, Newcombe VF, Simpson JP, Kane AD, Menon DK, Rueckert D, Glocker B. Efficient multi-scale 3D CNN with fully connected CRF for accurate brain lesion segmentation. *Medical image analysis*. 2017 Feb 1;36:61-78.
2. Havaei M, Davy A, Warde-Farley D, Biard A, Courville A, Bengio Y, Pal C, Jodoin PM, Larochelle H. Brain tumor segmentation with deep neural networks. *Medical image analysis*. 2017 Jan 1;35:18-31.

BLOCK

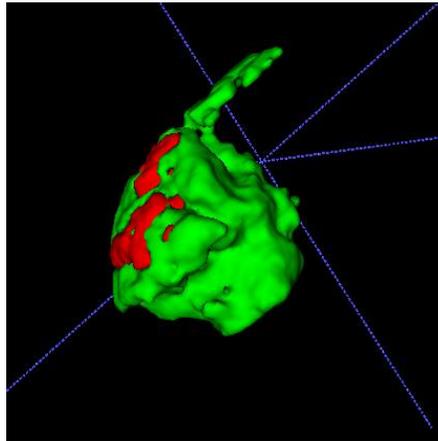


Block in the 3-D Network

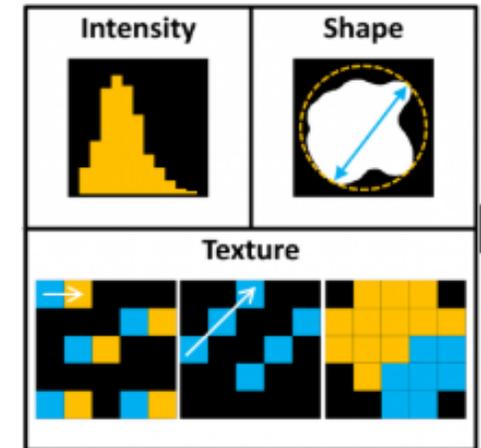
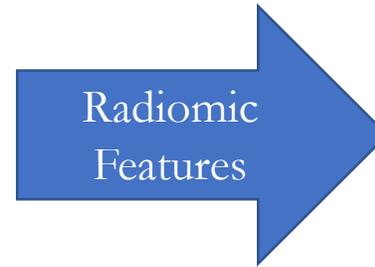
Radiomics Feature Extraction



Segmented T1 Image



3D Tumor Patch



Radiomic Feature Extraction¹

The features were extracted via the radiomics platform provided by pyradiomics². These features describe, amongst others, tumor image intensity, texture and shape and size of the tumor.

Shape Features (13)

First Order statistics (18)

GLCM Features (Gray Level Co-occurrence Matrix) (23)

GLRLM Features (Gray Level Run Length Matrix) (30)

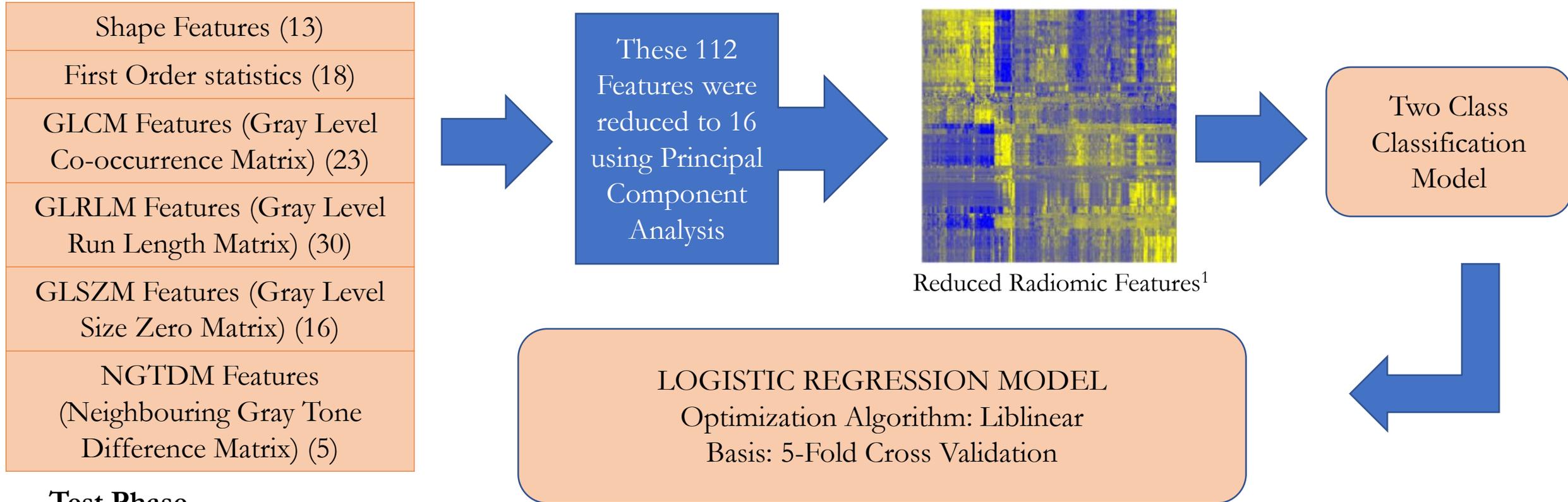
GLSZM Features (Gray Level Size Zero Matrix) (16)

NGTDM Features (Neighbouring Gray Tone Difference Matrix) (5)

1. <http://www.radiomics.world/?q=node/4>, 2. Joost JM van Griethuysen, Andriy Fedorov, Chintan Parmar, Ahmed Hosny, Nicole Aucoin, Vivek Narayan, Regina GH Beets-Tan, Jean-Christophe Fillion-Robin, Steve Pieper, and Hugo JWL Aerts. Computational radiomics system to decode the radiographic phenotype. Cancer research, 77(21):e104–e107, 2017.

Two Class Classification

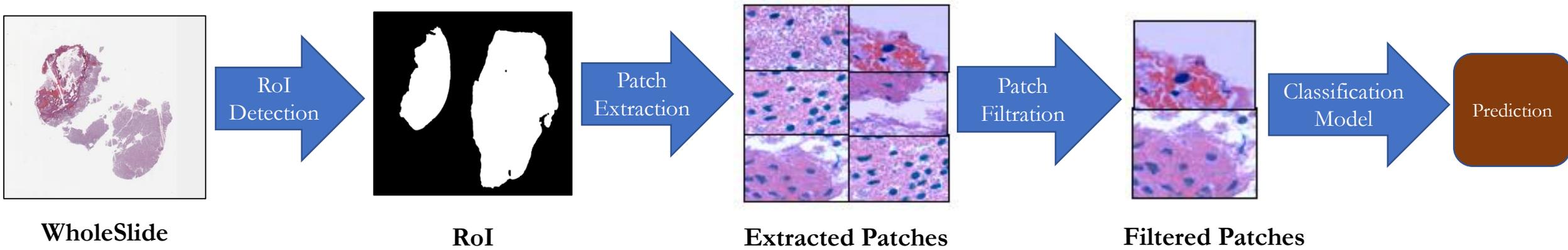
Each (16,1) length reduced radiomic features vector was used to train a Two Class Classification Model. Logistic Regression with LIBLINEAR as the optimization algorithm was used to train the model on a 5-fold cross validation basis.



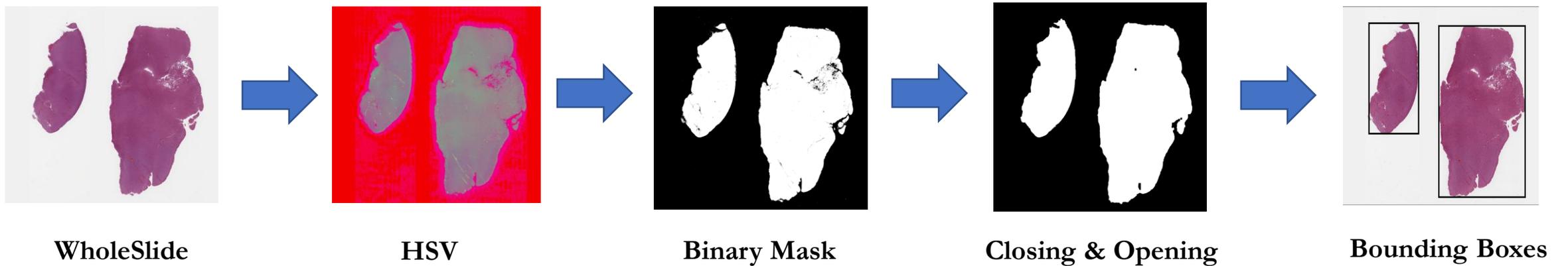
Test Phase

From each slide, patches are extracted exhaustively and stains are normalized. Trained autoencoder is used to extract features for each patch. Using these features outlier patches are filtered out using Isolation Forest. Voting based prediction on the set of filtered patches gives the class prediction

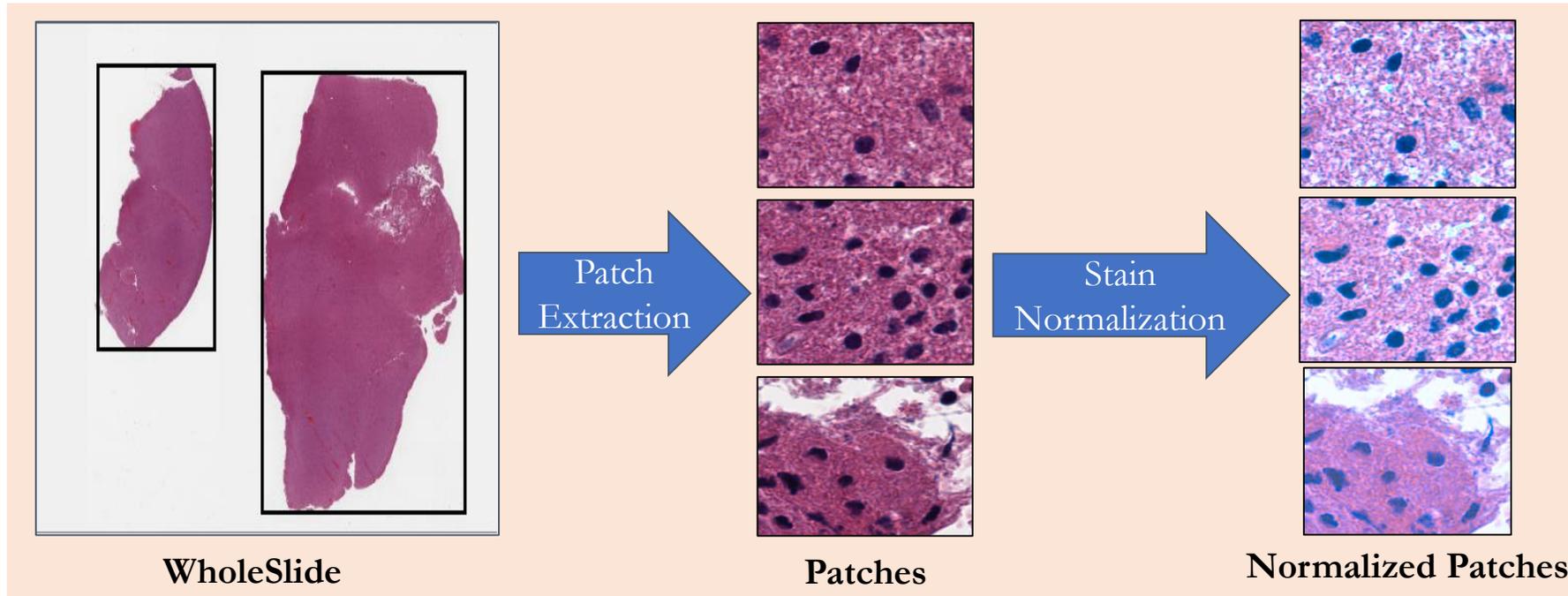
Histopathology Pipeline



- Region of Interest Detection



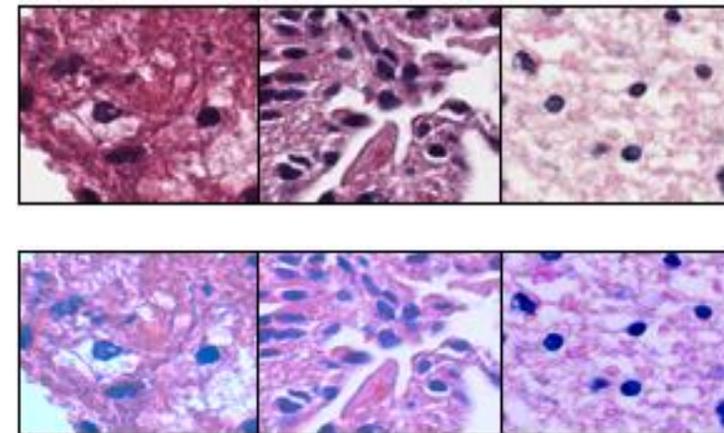
Patch Extraction and Stain Normalization



- Patches of size 224 x 224 were extracted from each Slide
- Patches extracted from Level-0 (highest resolution)
- Stain normalization¹ is used to normalize all the patches



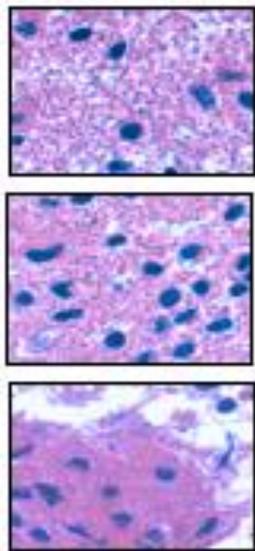
Variation in Stain Intensity



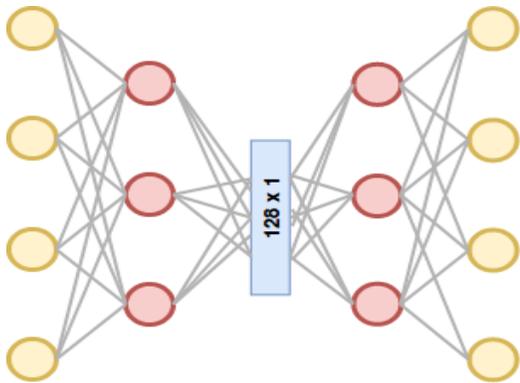
Before Normalization

After Normalization

Feature Extraction and Outlier Detection



Input



Convolutional Autoencoder

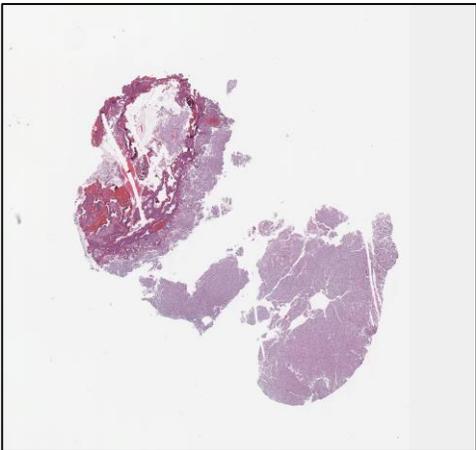
Output

Feature vector of size 128x1 for each patch

PCA

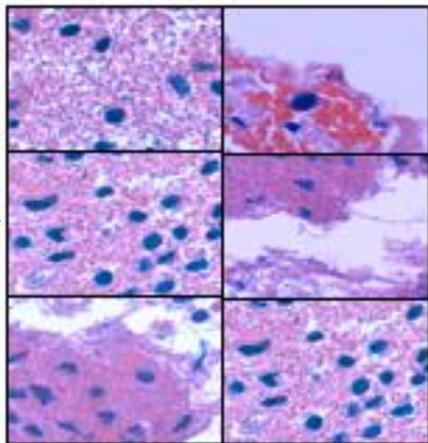
Dimensionality Reduced to 16x1

Why outlier detection? Outlier patches among all the patches in a pathology whole slide were used as possible differentials for two class classification.



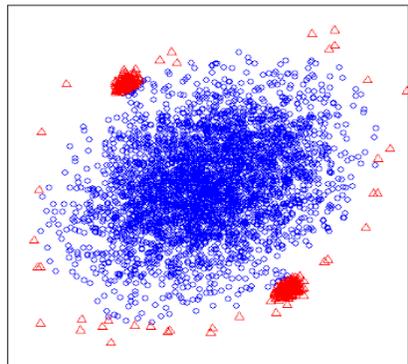
WholeSlide

Patches



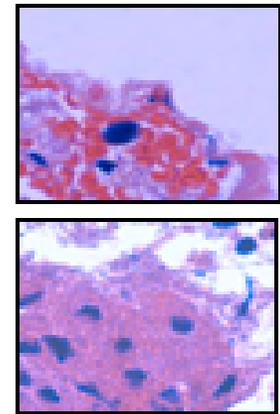
All patches from a single WholeSlide

Finding Outliers



Outlier Detection Using Isolation Forest

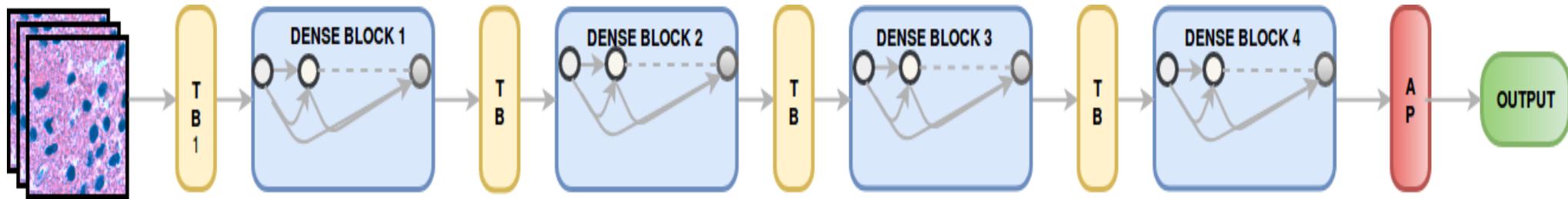
Outlier Patches



Training Data

Two Class Classification

- **Training Set:** 60k patches from 30 training slides used for training DenseNet Classifier
- Model trained using mini-batch Stochastic Gradient Descent

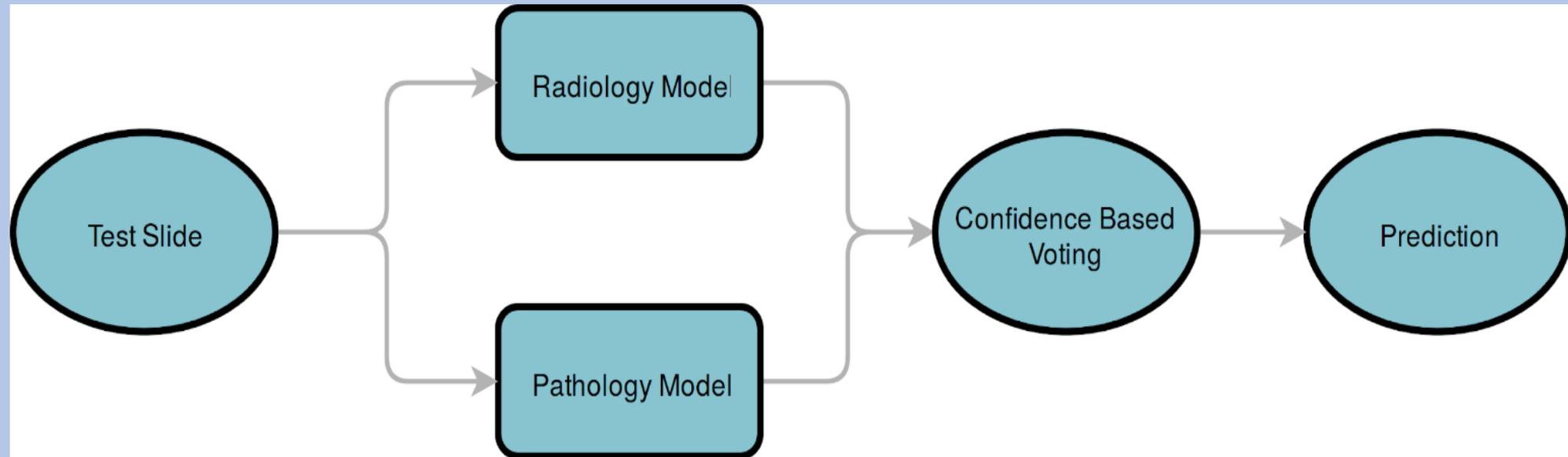


Test Phase

- From each slide, patches are extracted exhaustively and stains are normalized
- Trained autoencoder is used to extract features for each patch
- Using these features outlier patches are filtered out using Isolation Forest
- Voting based prediction on the set of filtered patches gives the class prediction

Combining Radiology with Pathology Results

- Both models achieve an accuracy of **80%** on the test set
- Confidence based voting was used to combine results from both the model
- Final accuracy with the combined results was **90%** on the test set



Pipeline for Combining Results from Radiology and Pathology