

# Improving Pathological Structure Segmentation Via Transfer Learning Across Diseases

**Barleen Kaur**, Paul Lemaitre, Raghav Mehta, Nazanin Mohammadi-Sepahvand,  
Doina Precup, Douglas L. Arnold and Tal Arbel

Workshop on Domain Adaptation and Representation Transfer, MICCAI 2019



# Motivation

- ❖ Lack of access to large annotated datasets: major challenges in medical imaging analysis.

# Motivation

- ❖ Lack of access to large annotated datasets: major challenges in medical imaging analysis.
- ❖ State-of-art models are based on deep learning methods, which perform well when trained on large datasets<sup>[1]</sup>.

[1] Özgün Çiçek et al, MICCAI 2016

[2] Veronika and et al., MIA 2019

# Motivation

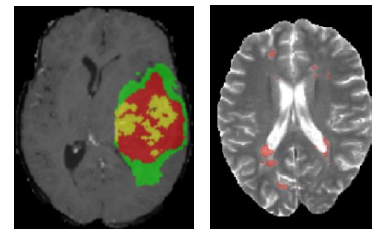
- ❖ Lack of access to large annotated datasets: major challenges in medical imaging analysis.
- ❖ State-of-art models are based on deep learning methods, which perform well when trained on large datasets<sup>[1]</sup>.
- ❖ Transfer learning has been explored in various applications such as classification, detection and segmentation. See <sup>[2]</sup> for a survey.

[1] Özgün Çiçek et al, MICCAI 2016

[2] Veronika and et al., MIA 2019

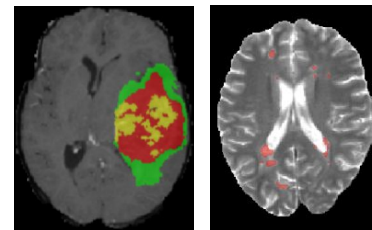
# Motivation

- ❖ Lack of access to large annotated datasets: major challenges in medical imaging analysis.
- ❖ State-of-art models are based on deep learning methods, which perform well when trained on large datasets<sup>[1]</sup>.
- ❖ Transfer learning has been explored in various applications such as classification, detection and segmentation. See <sup>[2]</sup> for a survey.
- ❖ Pathology segmentation:
  - Public datasets are small. Most of the large datasets are inhouse.
  - Difficult to obtain ground truth.
  - Class imbalance and inter-subject variability.

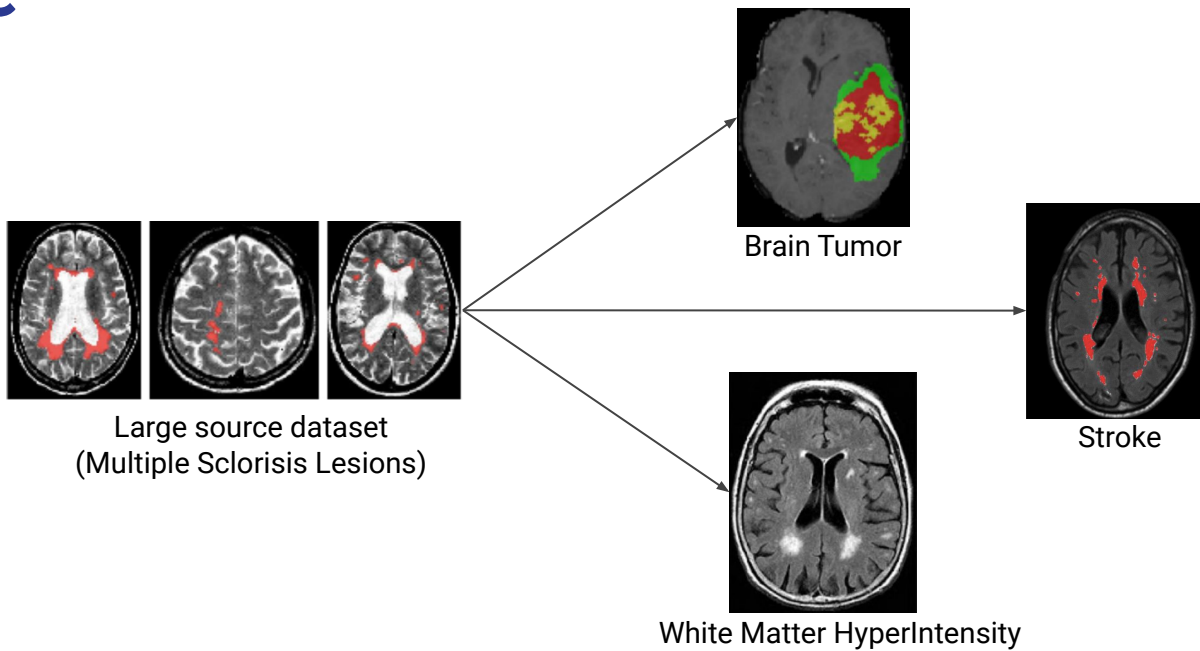


# Motivation

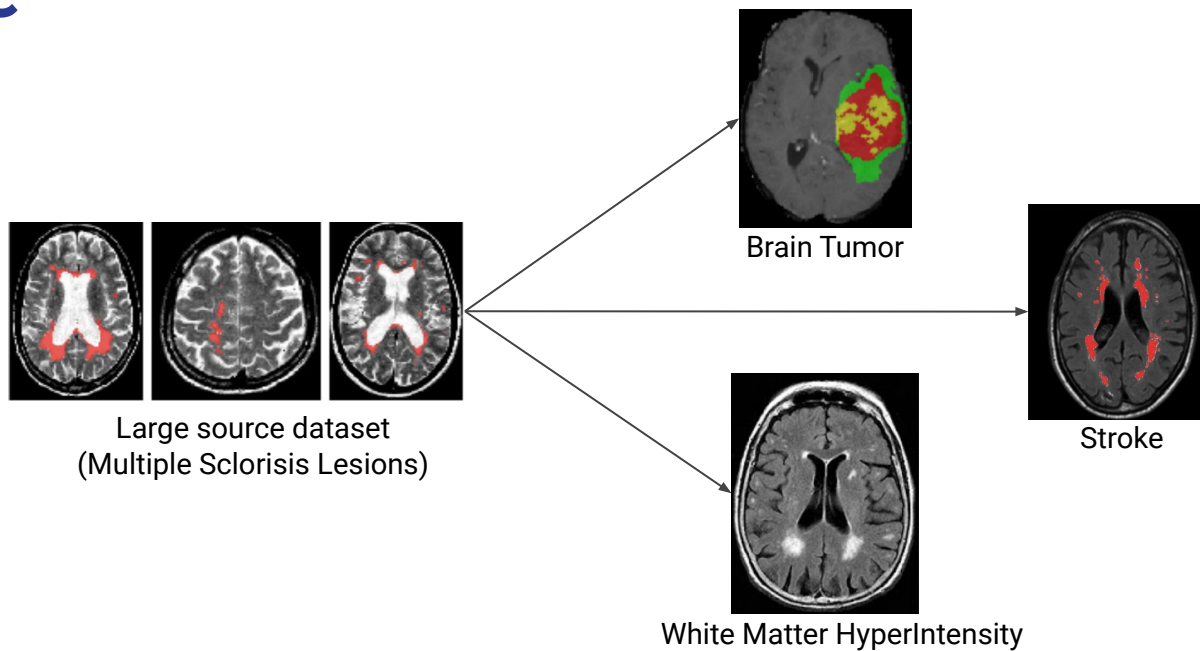
- ❖ Lack of access to large annotated datasets: major challenges in medical imaging analysis.
- ❖ State-of-art models are based on deep learning methods, which perform well when trained on large datasets<sup>[1]</sup>.
- ❖ Transfer learning has been explored in various applications such as classification, detection and segmentation. See <sup>[2]</sup> for a survey.
- ❖ Pathology segmentation:
  - Public datasets are small. Most of the large datasets are inhouse.
  - Difficult to obtain ground truth.
  - Class imbalance and inter-subject variability.
  - **Leveraging models trained on large datasets** in order to improve **pathology segmentation** results on smaller dataset **across different diseases** could be impactful in medical image analysis.



# Objective



# Objective

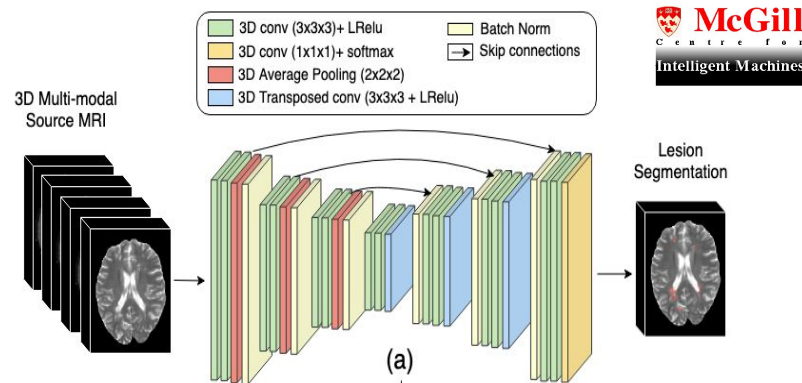


- ❖ Natural images: fine-tuning just last few layers helps. Is it same case in medical domain?
- ❖ We explore several fine-tuning strategies to see how to best leverage the source model and adapt it to the target dataset of varying sizes.



# Methodology

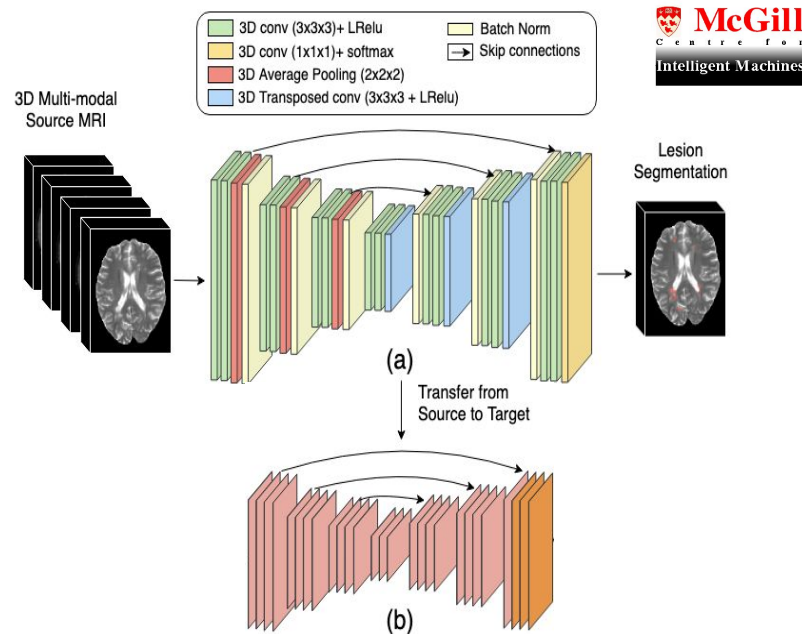
**First Phase:** Pretraining the UNet<sup>[1]</sup> with source MS dataset.



# Methodology

**First Phase:** Pretraining the UNet<sup>[1]</sup> with source MS dataset.

**Second Phase:** Replacing the last three task-specific layers of the pre-trained MS network with **new layers** and then fine-tuning with target brain tumor in three different ways:

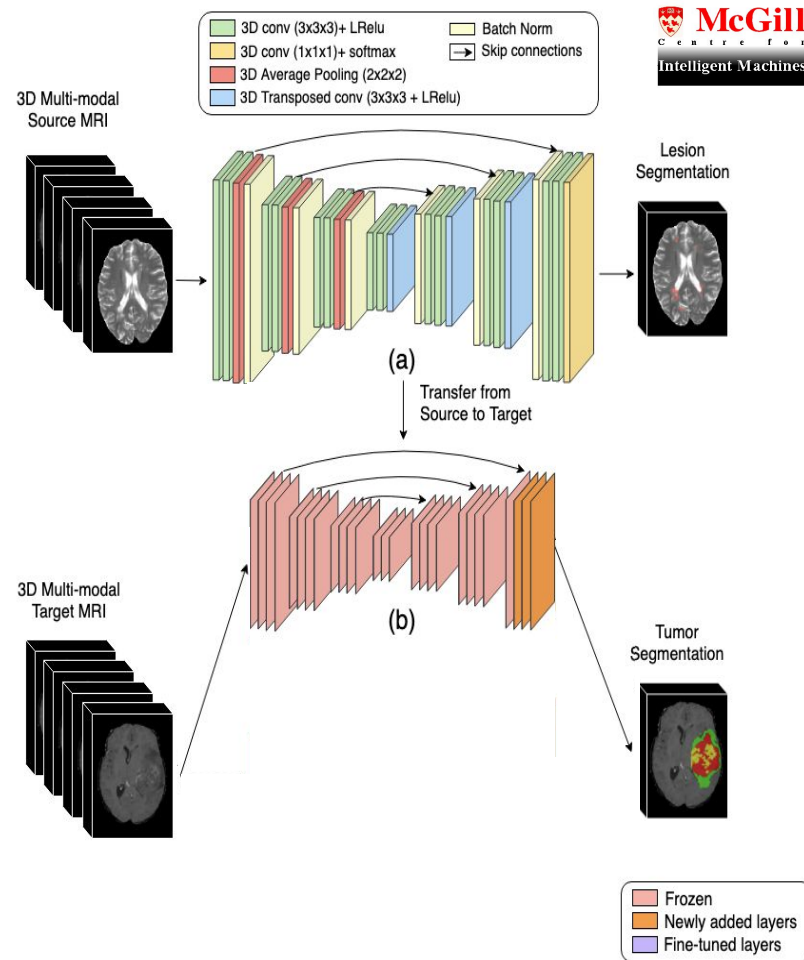


# Methodology

**First Phase:** Pretraining the UNet<sup>[1]</sup> with source MS dataset.

**Second Phase:** Replacing the last three task-specific layers of the pre-trained MS network with **new layers** and then fine-tuning with target brain tumor in three different ways:

- ★ **FT\_LastThree:** only the newly added layers are re-trained.

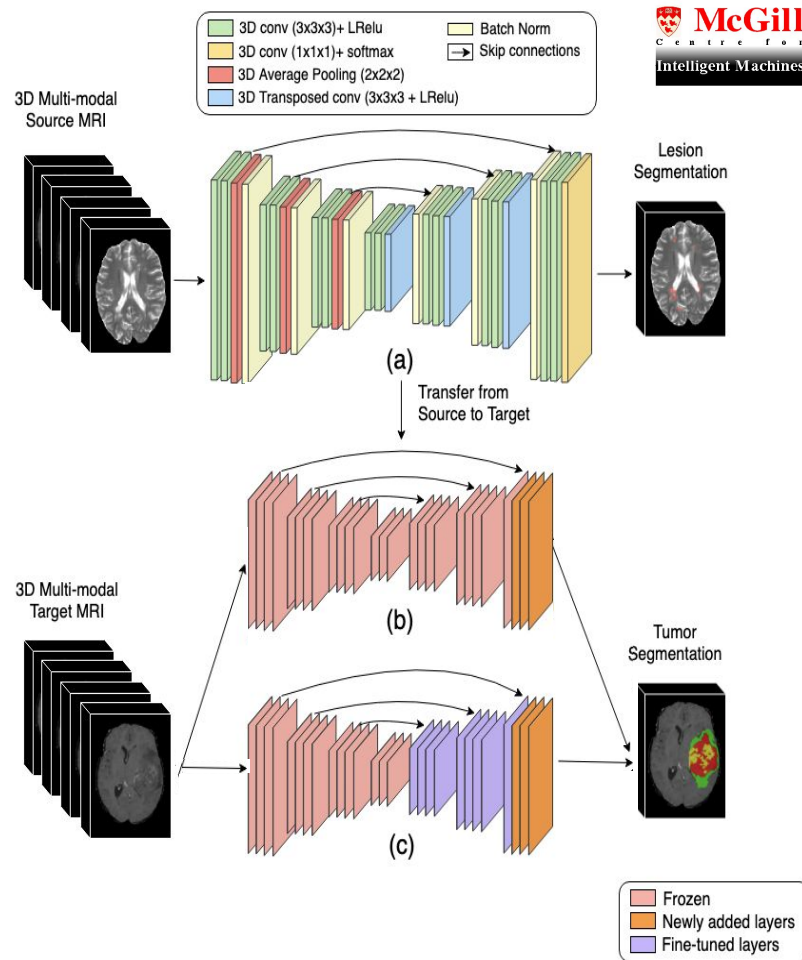


# Methodology

**First Phase:** Pretraining the UNet<sup>[1]</sup> with source MS dataset.

**Second Phase:** Replacing the last three task-specific layers of the pre-trained MS network with **new layers** and then fine-tuning with target brain tumor in three different ways:

- ★ **FT\_LastThree:** only the newly added layers are re-trained.
- ★ **FT\_Decoder:** Encoder part is frozen and only the decoder is fine-tuned.

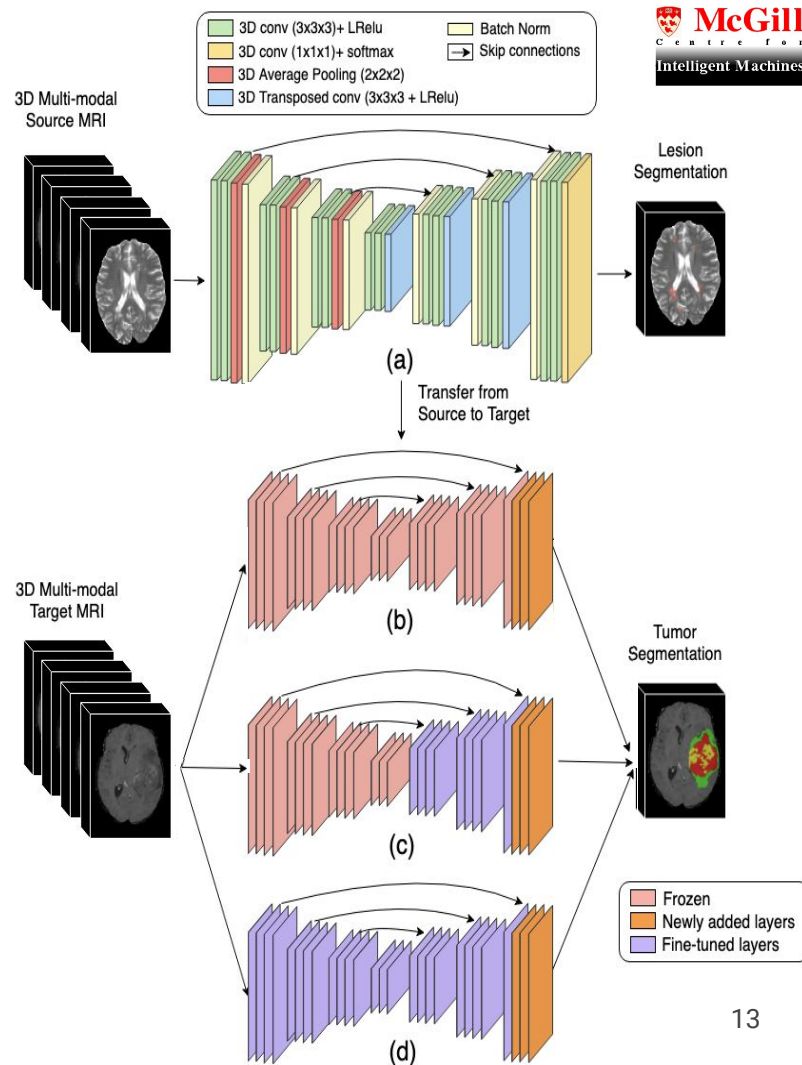


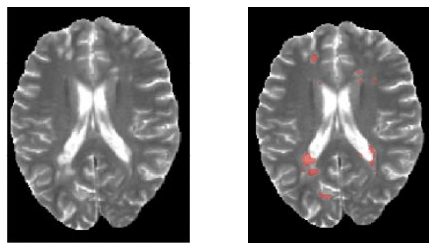
# Methodology

**First Phase:** Pretraining the UNet<sup>[1]</sup> with source MS dataset.

**Second Phase:** Replacing the last three task-specific layers of the pre-trained MS network with **new layers** and then fine-tuning with target brain tumor in three different ways:

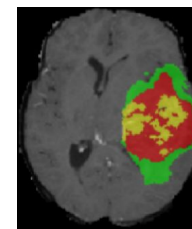
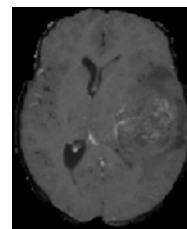
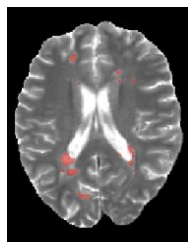
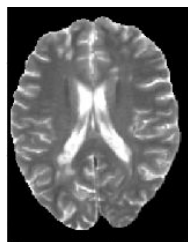
- ★ **FT\_LastThree:** only the newly added layers are re-trained.
- ★ **FT\_Decoder:** Encoder part is frozen and only the decoder is fine-tuned.
- ★ **FT\_All:** The whole pretrained network is fine-tuned.





## Source: Multiple Sclerosis Dataset

- ❖ Proprietary, multi-modal, multi-site, multi-scanner clinical trial dataset.
- ❖ 4 modalities (T1w, T2w, FLAIR, and T1 post-Gad)
- ❖ Resolution:  $1 \times 1 \times 1 \text{ mm}^3$
- ❖ Dimensions: 229x193x193.
- ❖ Total patient scans: 3630 multimodal MRI
- ❖ T2 binary lesion segmentation mask provided.



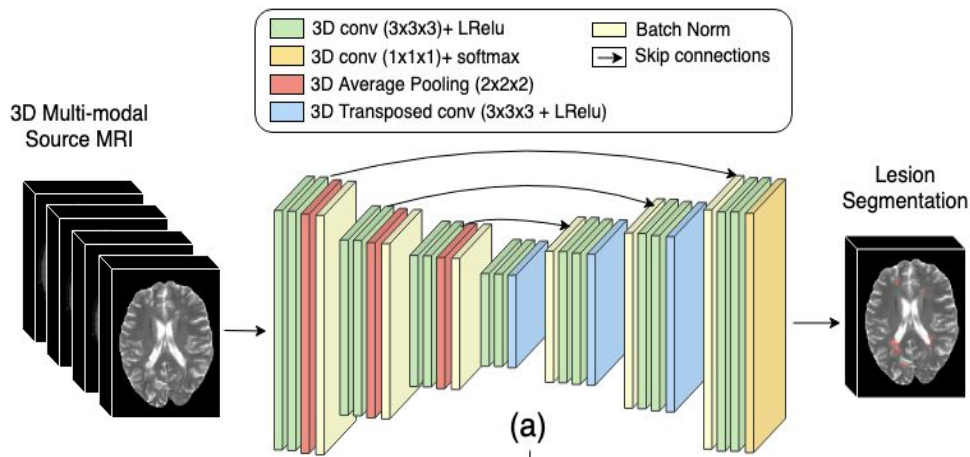
## Source: Multiple Sclerosis Dataset

- ❖ Proprietary, multi-modal, multi-site, multi-scanner clinical trial dataset.
- ❖ 4 modalities (T1w, T2w, FLAIR, and T1 post-Gad)
- ❖ Resolution:  $1 \times 1 \times 1 \text{ mm}^3$
- ❖ Dimensions:  $229 \times 193 \times 193$ .
- ❖ Total patient scans: 3630 multimodal MRI
- ❖ T2 binary lesion segmentation mask provided.

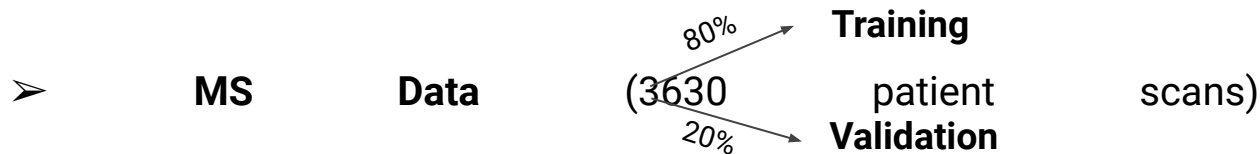
## Target: BraTS 2018 challenge Dataset<sup>[1]</sup>

- ❖ 4 modalities (T1, T2, FLAIR, T1c)
- ❖ Resolution:  $1 \times 1 \times 1 \text{ mm}^3$
- ❖ Dimensions:  $155 \times 240 \times 240$
- ❖ Manual marking for 3 types of tumor (edema, necrotic core, and enhancing core)
- ❖ BraTS 2018 Training data (285 patients) for training (Ground Truth available)
- ❖ BraTS 2018 Validation data (66 patients) for testing (Ground truth not provided)

# Experimentation (First Phase: Pre-training)

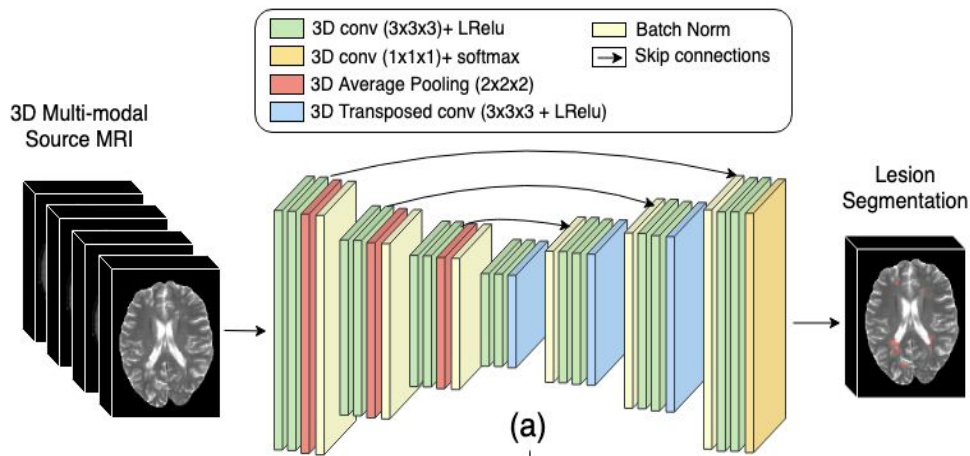


Pre-training the UNet with source MS data for T2 lesion segmentation.

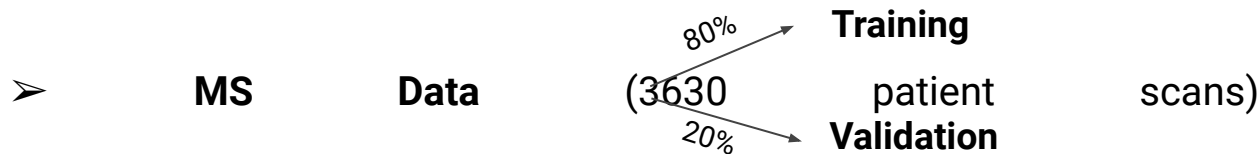




# Experimentation (First Phase: Pre-training)

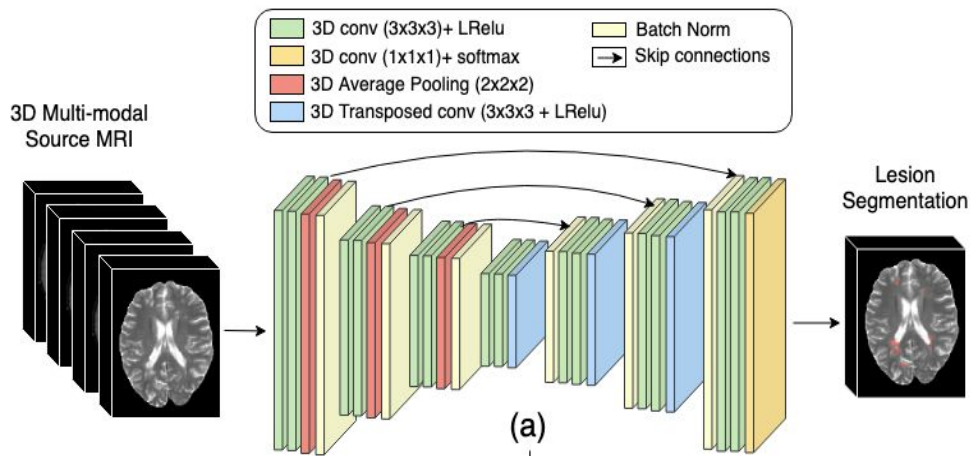


Pre-training the UNet with source MS data for T2 lesion segmentation.

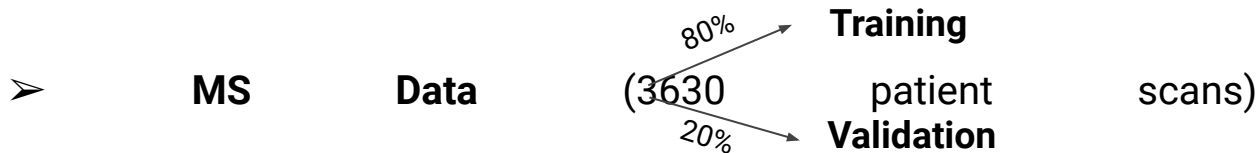


- Weighted binary cross entropy was used as loss function.

# Experimentation (First Phase: Pre-training)



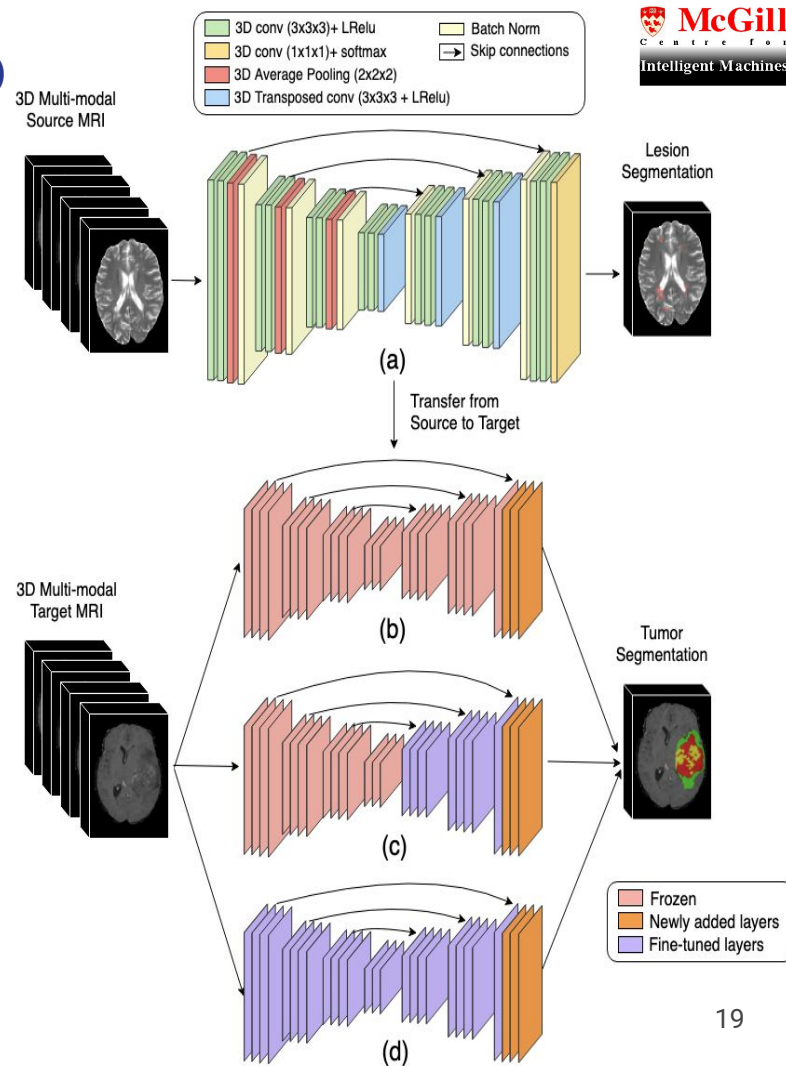
Pre-training the UNet with source MS data for T2 lesion segmentation.



- Weighted binary cross entropy was used as loss function.
- An **AUC of 0.77** was obtained on the validation (test) set.

# Experimentation (Second Phase: Fine-tuning)

For 20, 50, 100, 150 brain tumor MRI scans:

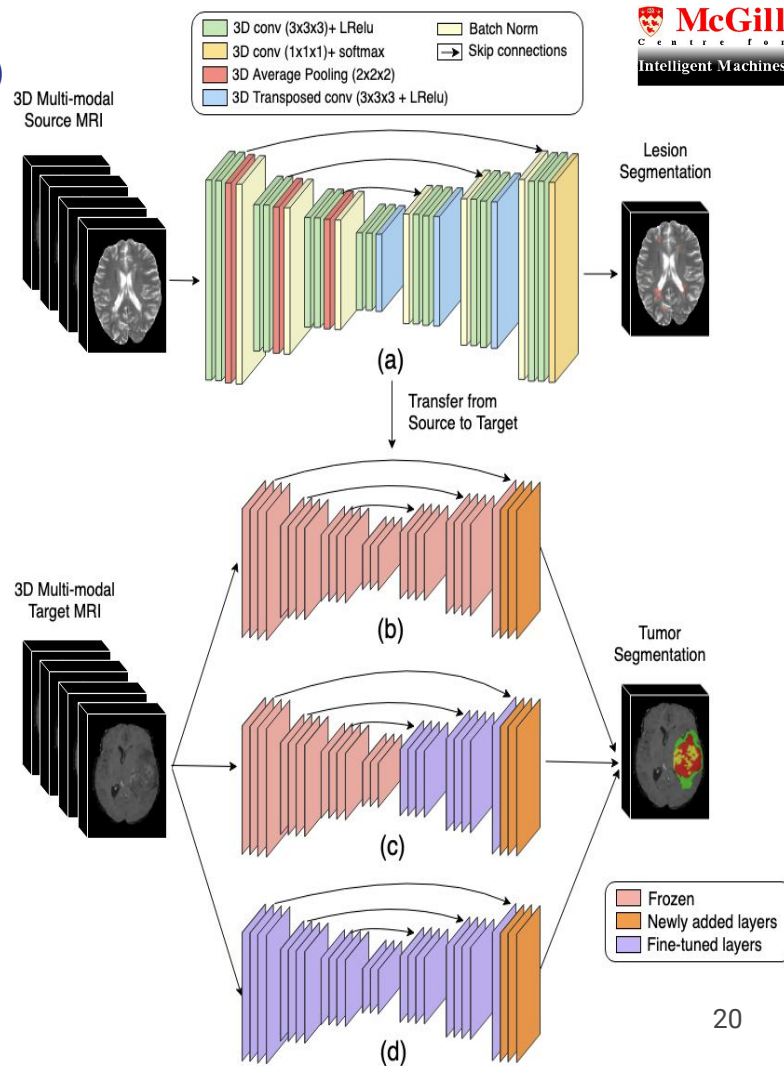


# Experimentation (Second Phase: Fine-tuning)

For 20, 50, 100, 150 brain tumor MRI scans:

## ★ Transfer Learning

- FT\_Last Three
- FT\_Decoder
- FT\_All



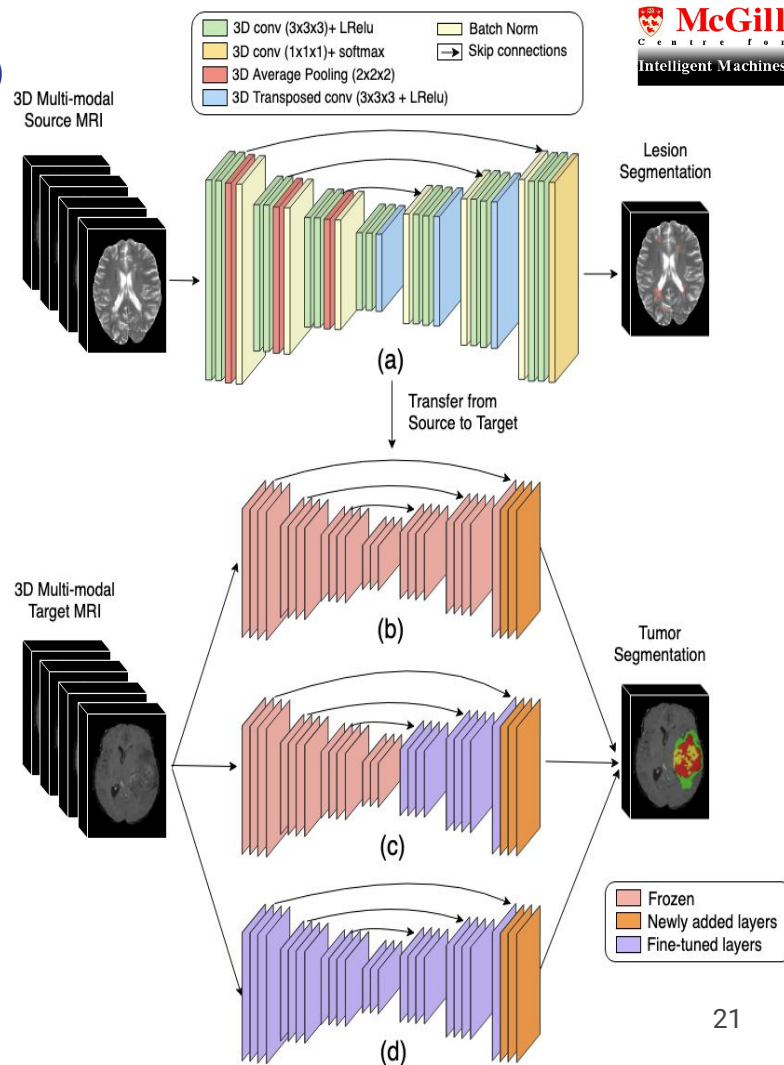
# Experimentation (Second Phase: Fine-tuning)

For 20, 50, 100, 150 brain tumor MRI scans:

## ★ Transfer Learning

- FT\_Last Three
- FT\_Decoder
- FT\_All

## ★ Baseline (Training from scratch)



# Experimentation (Second Phase: Fine-tuning)

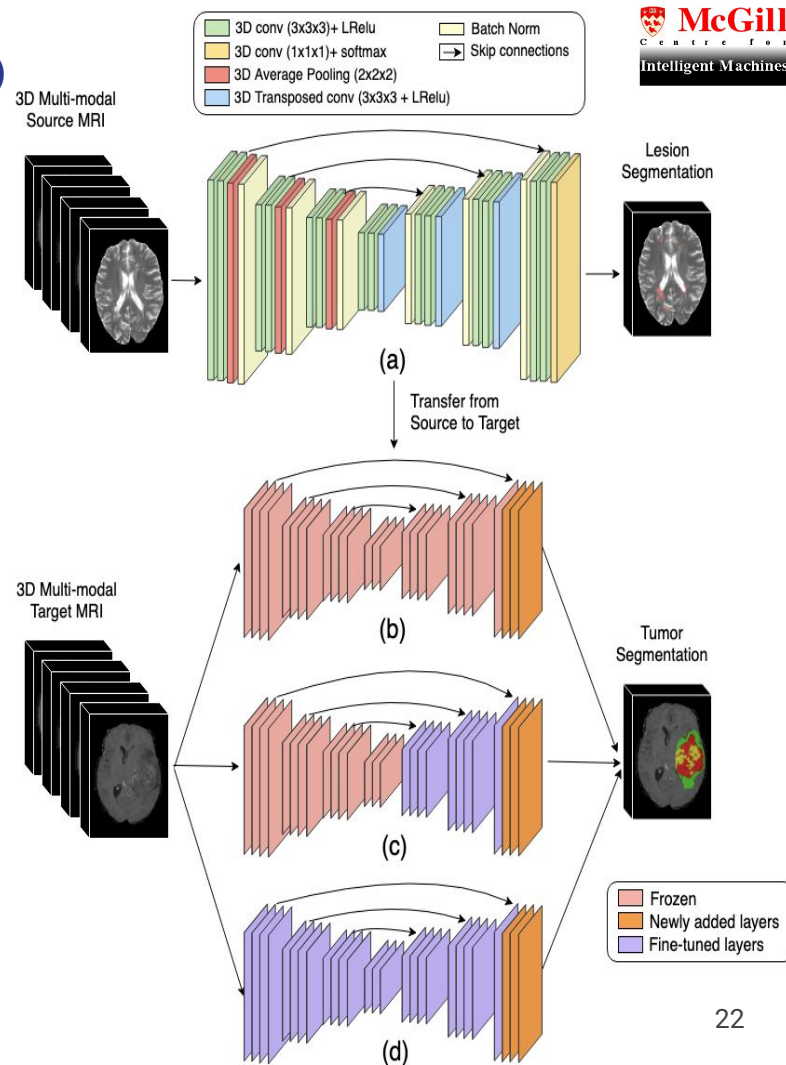
For 20, 50, 100, 150 brain tumor MRI scans:

## ★ Transfer Learning

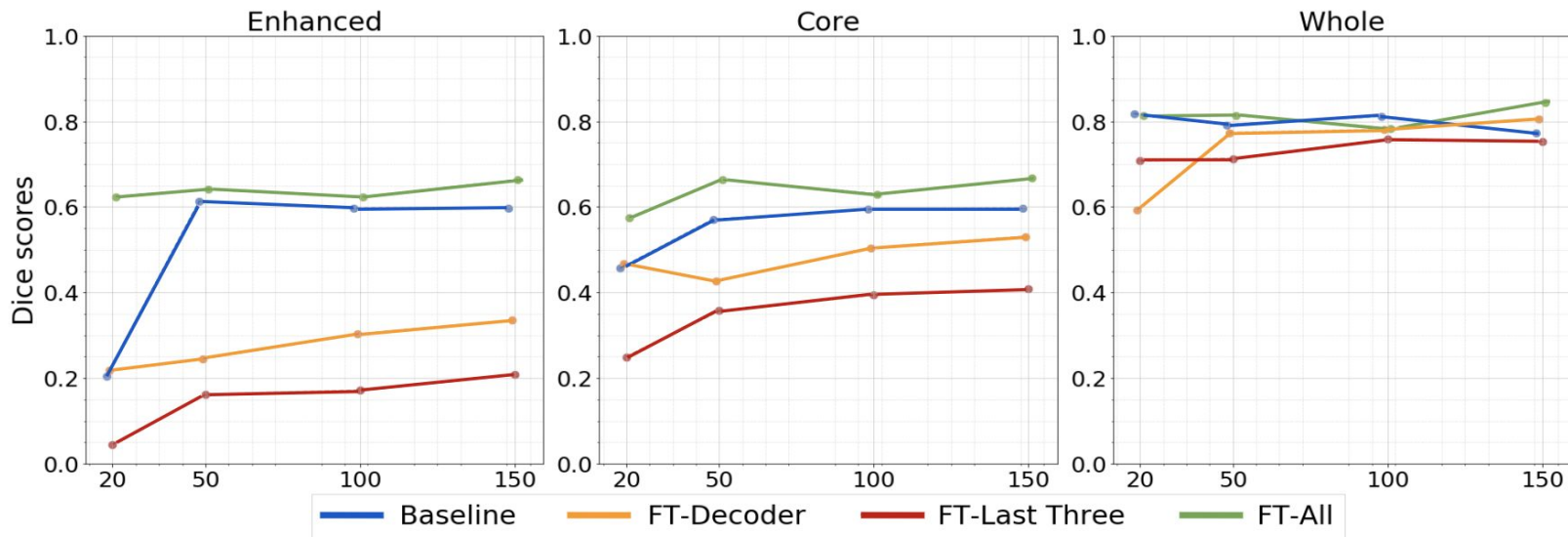
- FT\_Last Three
- FT\_Decoder
- FT\_All

## ★ Baseline (Training from scratch)

- Weighted Cross entropy loss.
- Four-fold cross validation
- A **local validation set** of 50 samples is used to select the operating point.

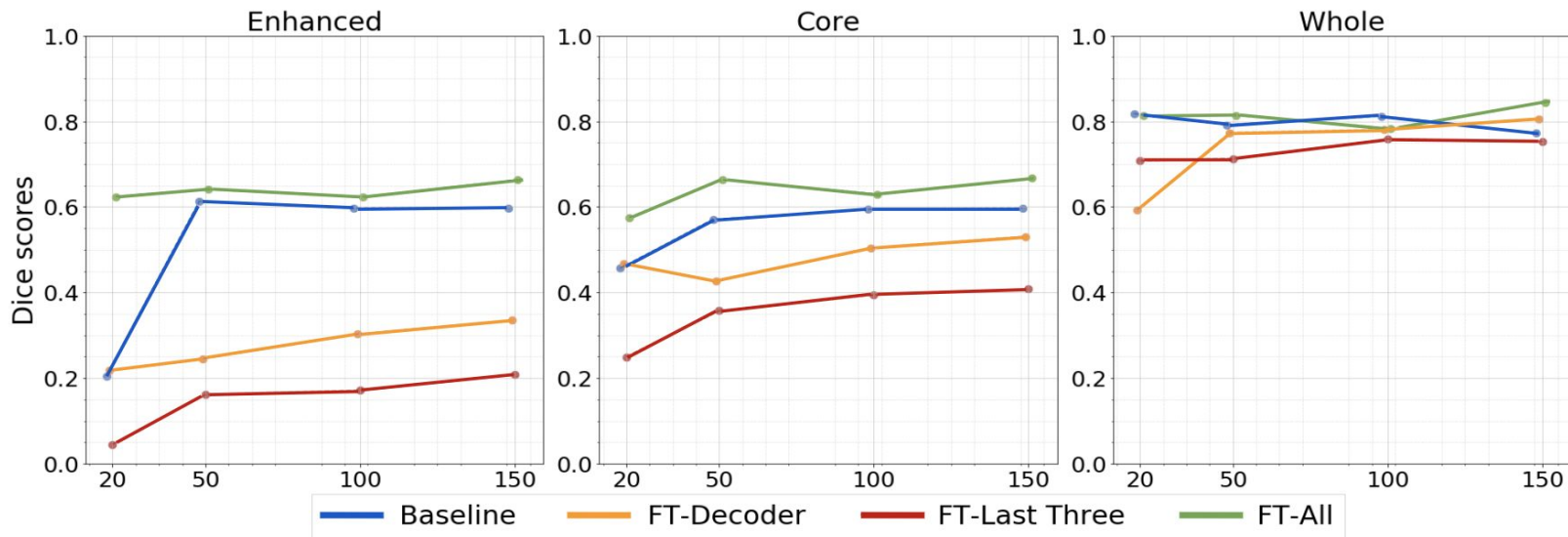


# Quantitative Results (on BraTS 2018 Validation set)



➤ **FT-All outperforms the baseline** in almost every case.

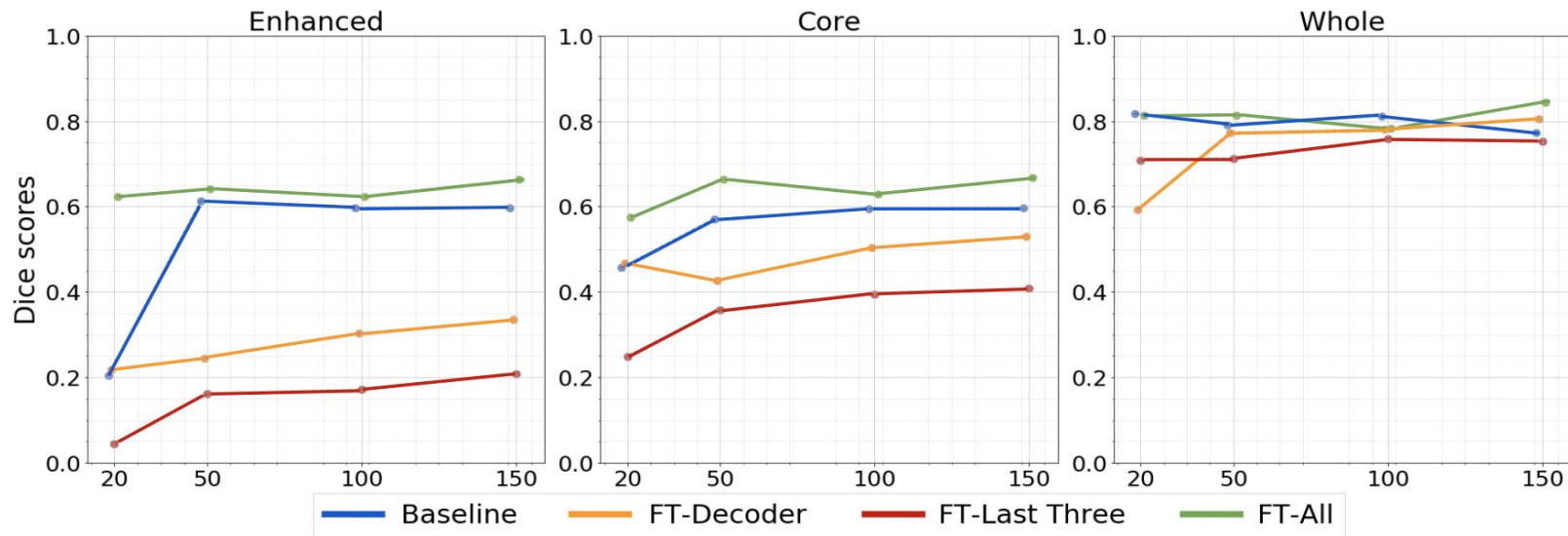
# Quantitative Results (on BraTS 2018 Validation set)



- **FT-All outperforms the baseline** in almost every case.
- Best when the **number of tumor cases is extremely low**, i.e. 20.



# Quantitative Results (on BraTS 2018 Validation set)

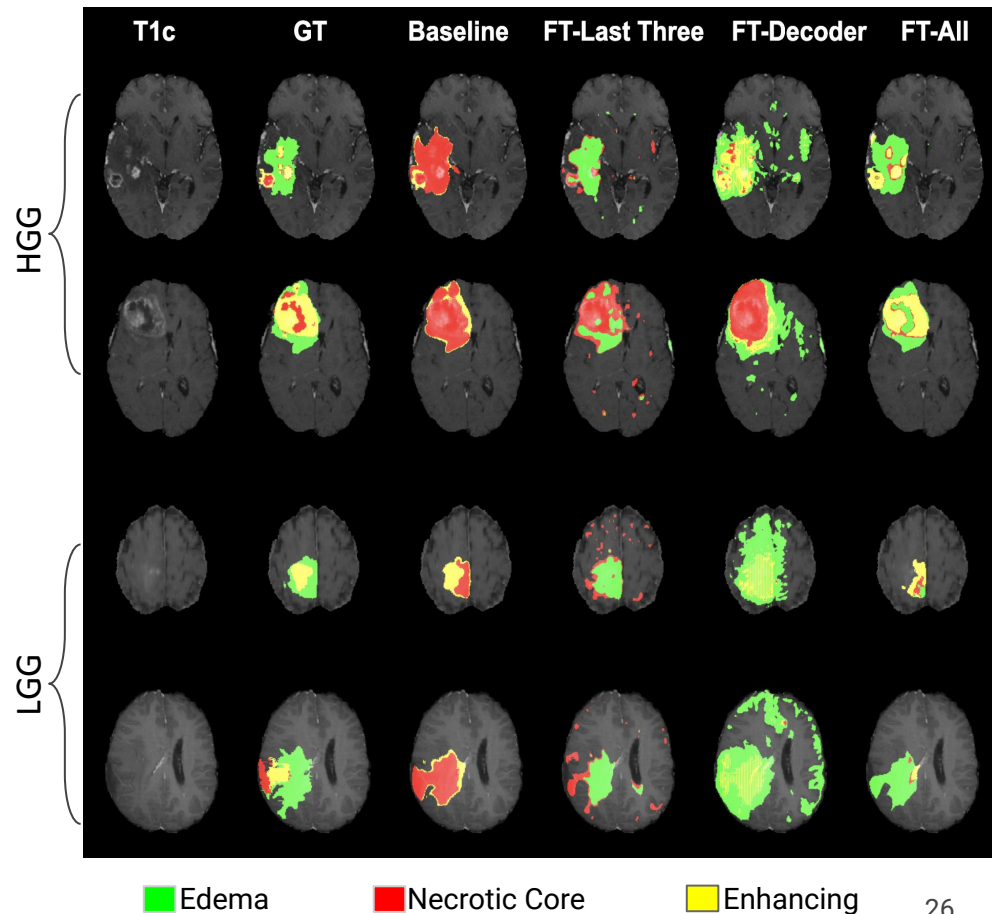


- **FT-All outperforms the baseline** in almost every case,
- Best when the **number of tumor cases is extremely low**, i.e. 20.
- As the number of brain tumor samples increase, **the gain of FT-All over baseline diminishes**.

# Qualitative Results

Fine-tuned with 20 brain tumor cases

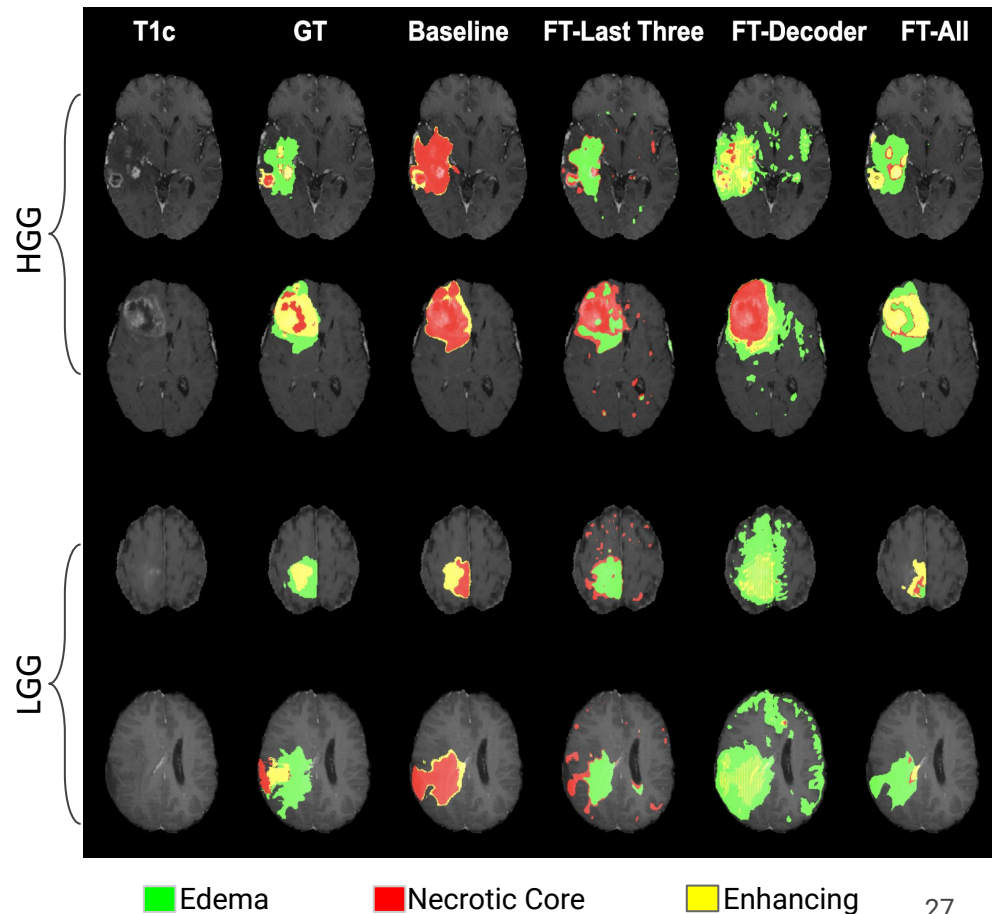
- FT-All is able to capture sub-structures of tumor better than the other methods.



# Qualitative Results

Fine-tuned with 20 brain tumor cases

- **FT-All is able to capture sub-structures of tumor better than the other methods.**
- Performance is better on the HGG over the LGG cases, as more HGG cases are present in the training dataset.



# Conclusions and Discussions

- ❖ We explored different strategies for transfer learning across diseases for the task of focal pathology segmentation.

# Conclusions and Discussions

- ❖ We explored different strategies for transfer learning across diseases for the task of focal pathology segmentation.
- ❖ We observed that **fine-tuning the whole network** works best, especially when **very small target datasets are available**.

# Conclusions and Discussions

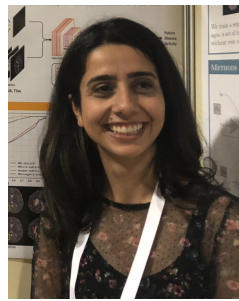
- ❖ We explored different strategies for transfer learning across diseases for the task of focal pathology segmentation.
- ❖ We observed that **fine-tuning the whole network** works best, especially when **very small target datasets are available**.
- ❖ We also observed that as in case of natural images, where fine-tuning just the last few layers works, **it's not the same case in medical domain**.

# Conclusions and Discussions

- ❖ We explored different strategies for transfer learning across diseases for the task of focal pathology segmentation.
- ❖ We observed that **fine-tuning the whole network** works best, especially when **very small target datasets are available**.
- ❖ We also observed that as in case of natural images, where fine-tuning just the last few layers works, **it's not the same case in medical domain**.
- ❖ We motivate **public release of models** trained on large datasets.

# Acknowledgement

- ❖ Lab mates and supervisors: Prof Tal Arbel and Prof Doina Precup.



- ❖ Sponsors



INTERNATIONAL  
PROGRESSIVE MS ALLIANCE

**CONNECT** TO END PROGRESSIVE MS



**Thank you for your patient listening!**

**Questions??**

## Source: Multiple Sclerosis Dataset

- ❖ Brain extraction<sup>[2]</sup>
- ❖ N3 bias field inhomogeneity correction<sup>[3]</sup>
- ❖ Nyul image intensity normalization<sup>[4]</sup>
- ❖ Registration to the MNI-space.
- ❖ Intensity Normalization (mean subtraction, divide by standard deviation, re-mapping to 0-1)
- ❖ Cropped and zero-padded to 240x192x192.

## Target: BraTS 2018 challenge Dataset <sup>[1]</sup>

- ❖ Skull stripping
- ❖ Co-registration
- ❖ Registration to same space as source data using ANTs tool<sup>[5]</sup>
- ❖ Intensity Normalization (mean subtraction, divide by standard deviation, re-mapping to 0-1)
- ❖ Cropped and zero-padded to 240x192x192.

[1] Menze et al, TMI 2015

[2] Smith et al, HBM 2002

[3] Sled et al TMI 1998

[4] Nyul et al TMI 2000

[5] Avants et al Neuroimage 2011