

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^[1].

[1] Özgün Çiçek et al, MICCAI 2016

[2] Veronika and et al., MIA 2019

Motivation

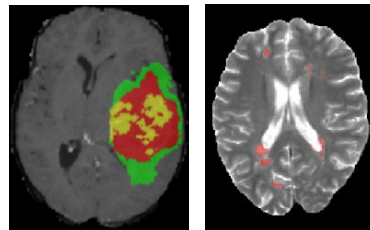
- ❖ Lack of access to large annotated datasets: major challenges in medical imaging analysis.
- ❖ State-of-art models are based on deep learning methods, which perform well when trained on large datasets^[1].
- ❖ Transfer learning has been explored in various applications such as classification, detection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

[1] Özgün Çiçek et al, MICCAI 2016

[2] Veronika and et al., MIA 2019

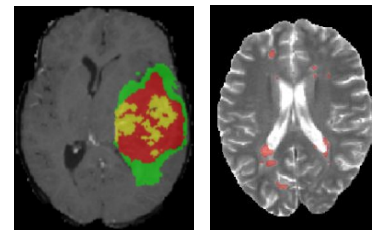
Motivation

- ❖ 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^[1].
- ❖ Transfer learning has been explored in various applications such as classification, detection and segmentation. See ^[2] 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.

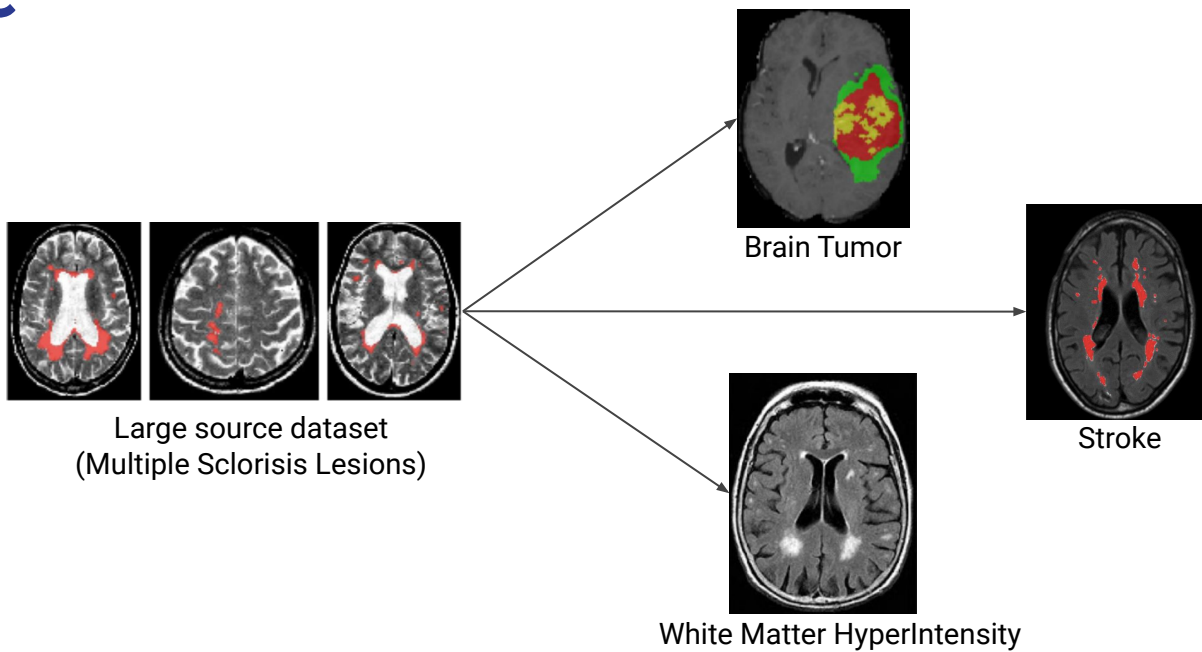


Motivation

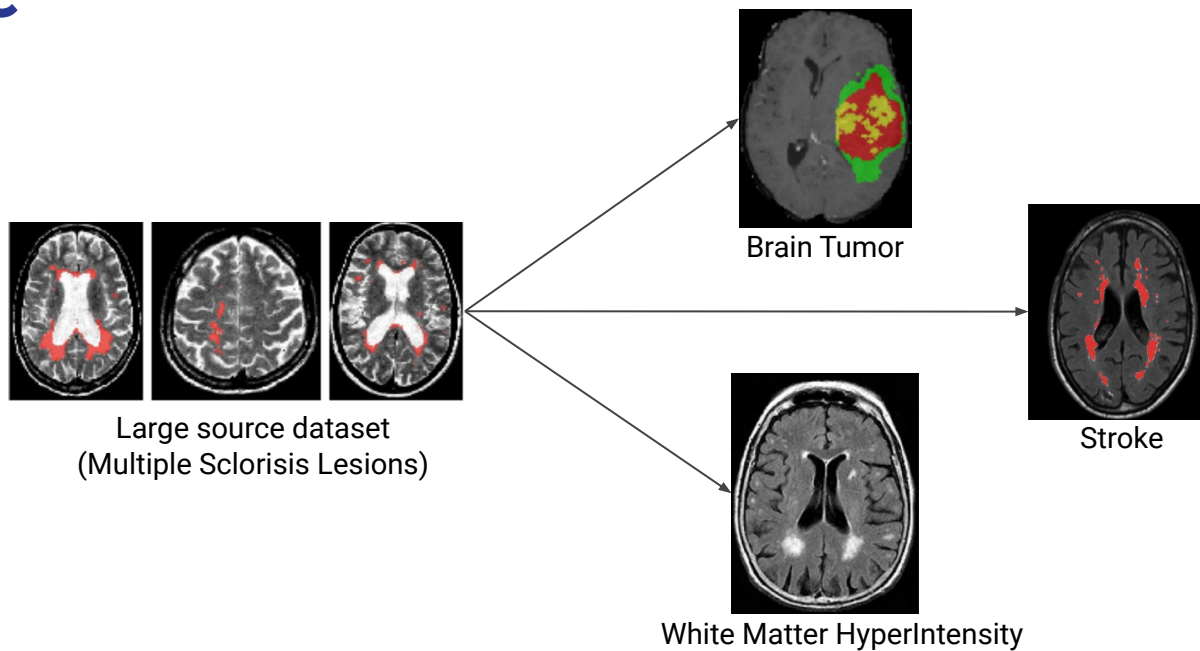
- ❖ Lack of access to large annotated datasets: major challenges in medical imaging analysis.
- ❖ State-of-art models are based on deep learning methods, which perform well when trained on large datasets^[1].
- ❖ Transfer learning has been explored in various applications such as classification, detection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



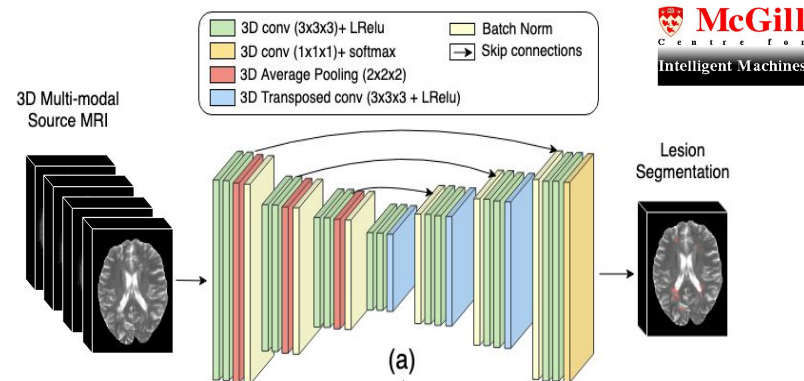
Objective



- ❖ 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

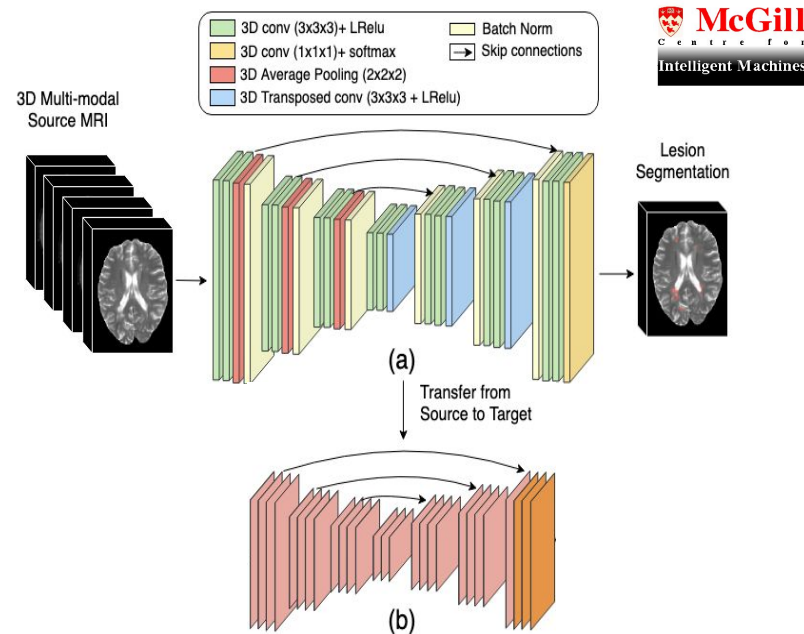
First Phase: Pretraining the UNet^[1] with source MS dataset.



Methodology

First Phase: Pretraining the UNet^[1] 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:

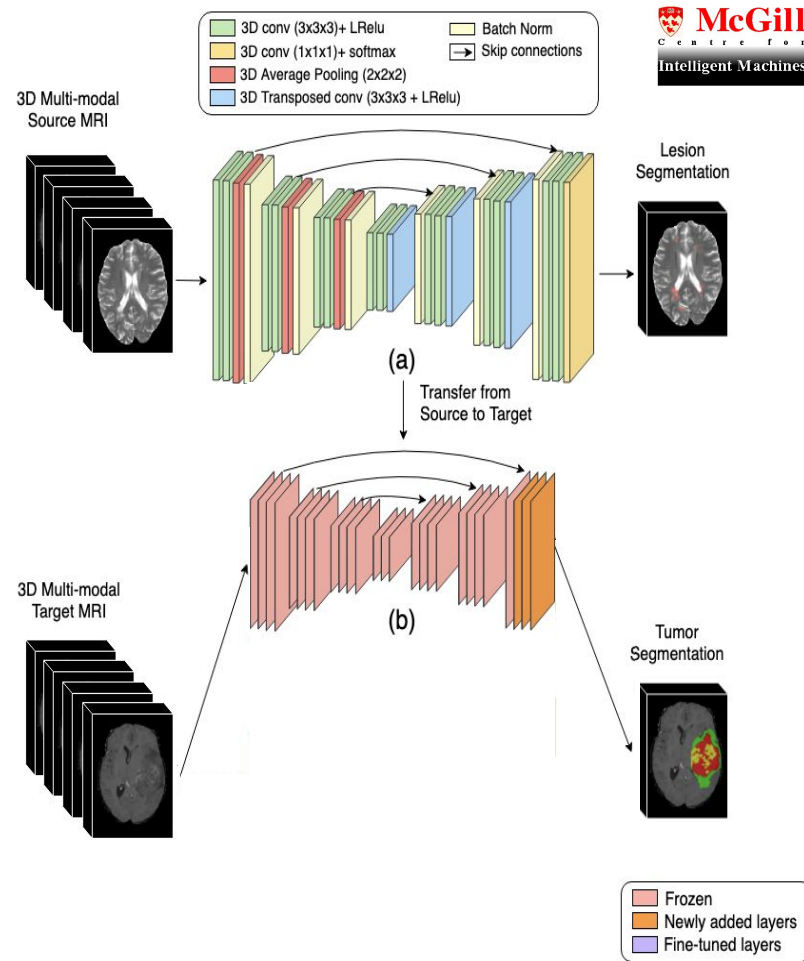


Methodology

First Phase: Pretraining the UNet^[1] 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.

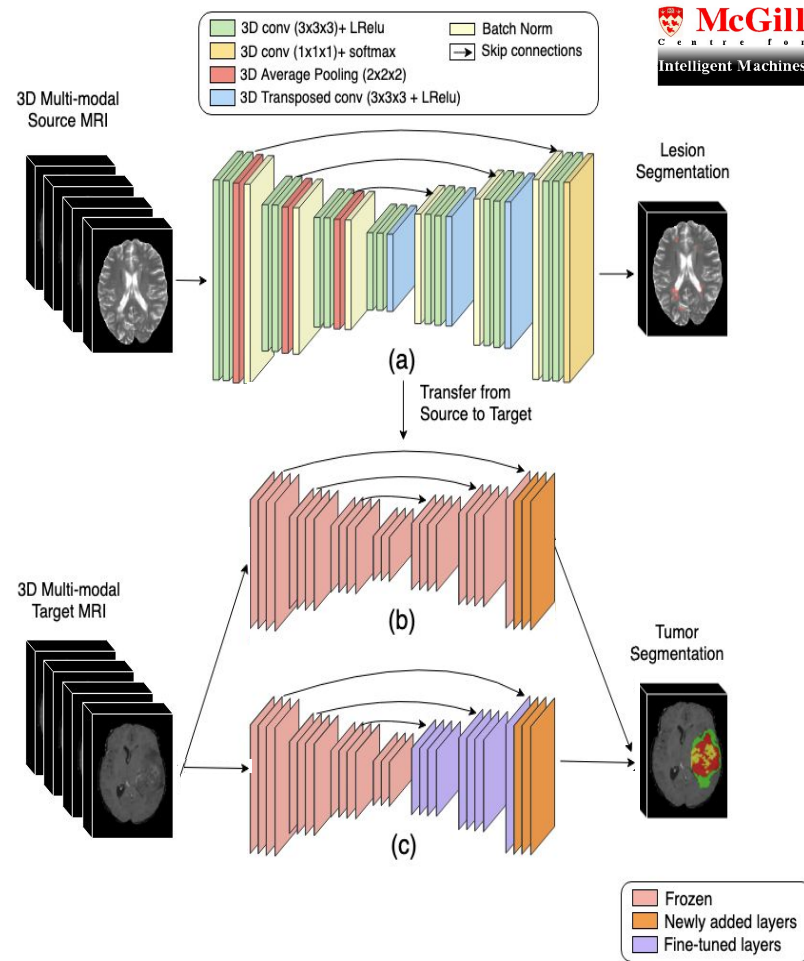


Methodology

First Phase: Pretraining the UNet^[1] 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.

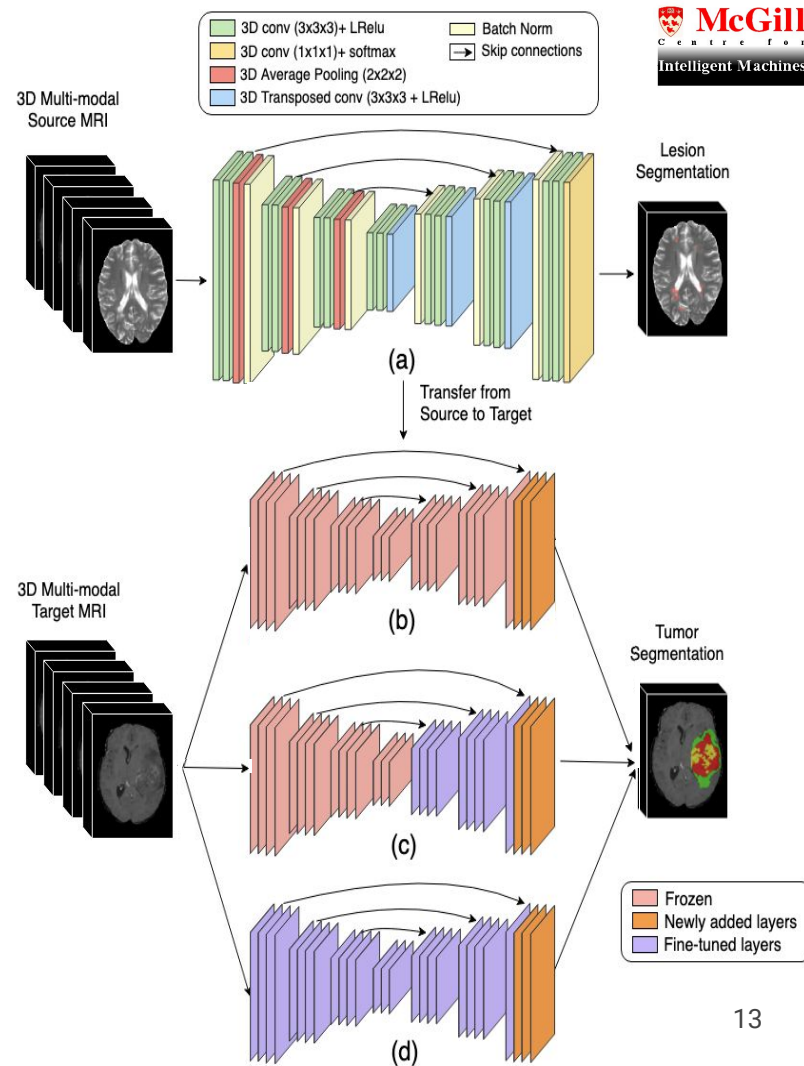


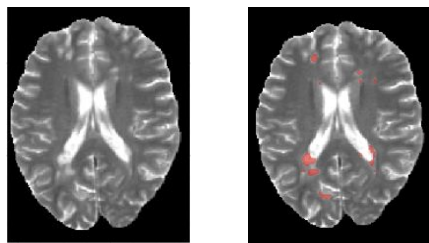
Methodology

First Phase: Pretraining the UNet^[1] 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:

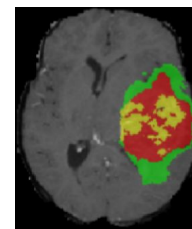
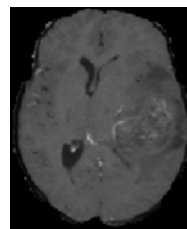
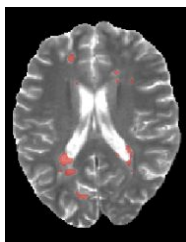
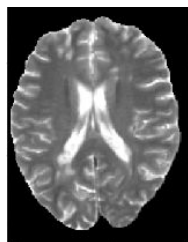
- ★ **FT_LastThree:** only the newly added layers are re-trained.
- ★ **FT_Decoder:** Encoder part is frozen and only the decoder is fine-tuned.
- ★ **FT_All:** The whole pretrained network is fine-tuned.





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: $229 \times 193 \times 193$.
- ❖ Total patient scans: 3630 multimodal MRI
- ❖ T2 binary lesion segmentation mask provided.



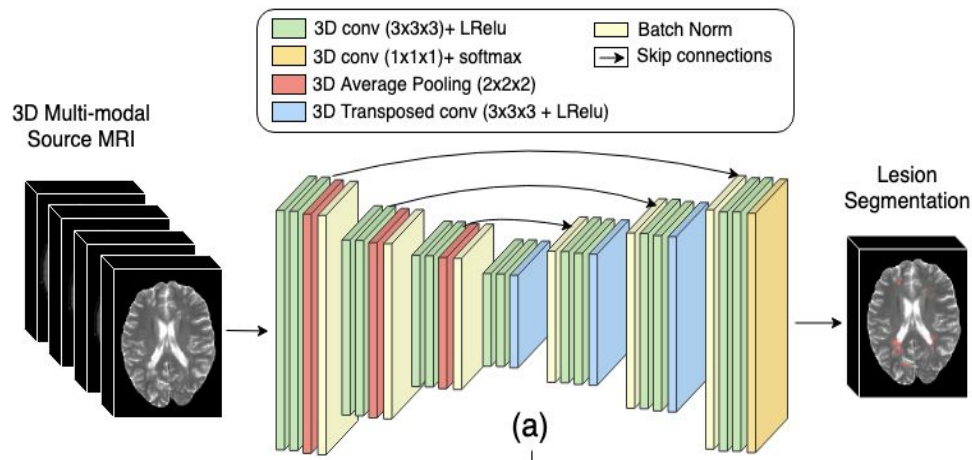
Source: Multiple Sclerosis Dataset

- ❖ 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.

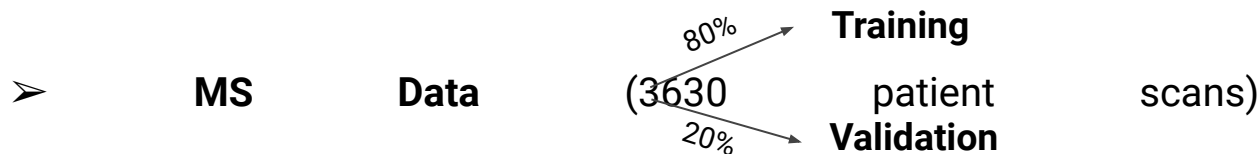
Target: BraTS 2018 challenge Dataset^[1]

- ❖ 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)

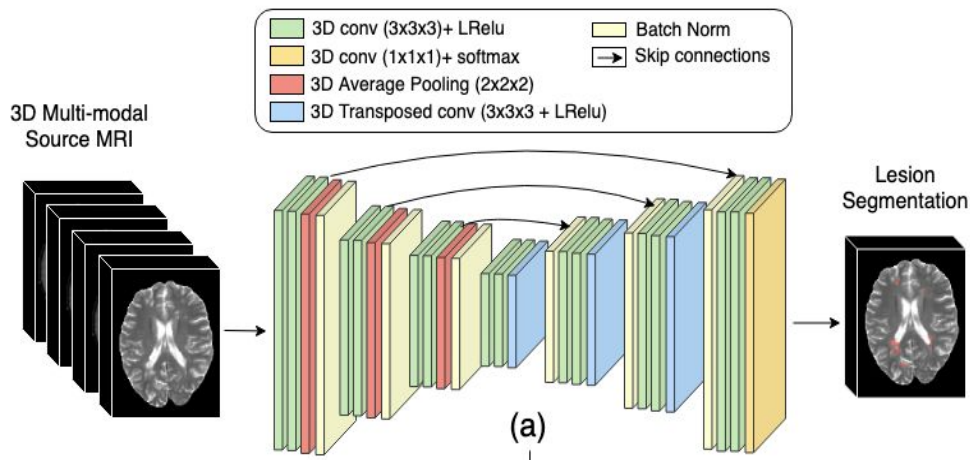
Experimentation (First Phase: Pre-training)



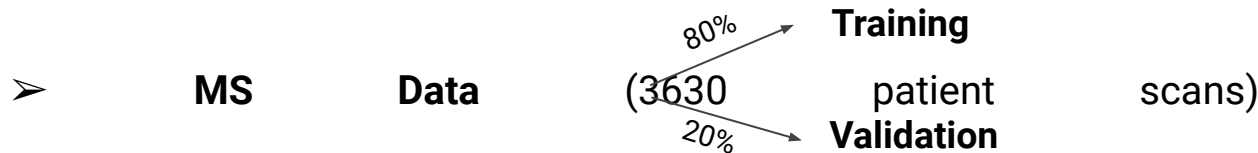
Pre-training the UNet with source MS data for T2 lesion segmentation.



Experimentation (First Phase: Pre-training)

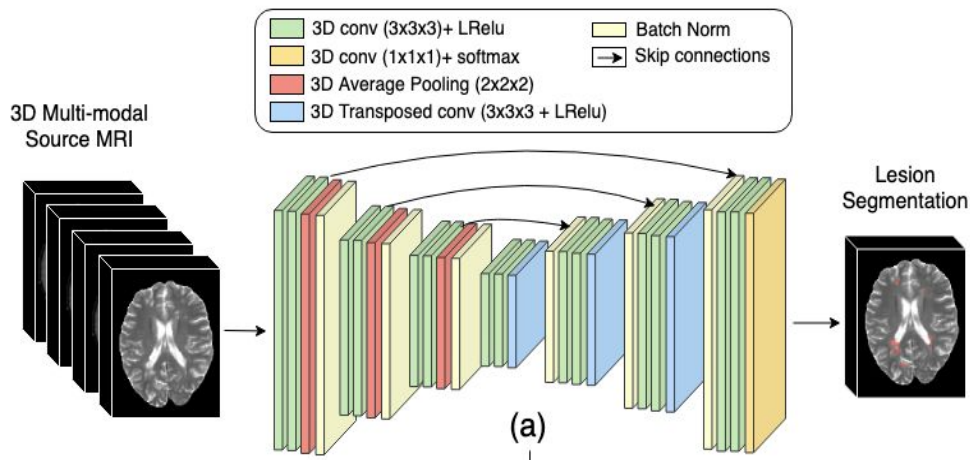


Pre-training the UNet with source MS data for T2 lesion segmentation.

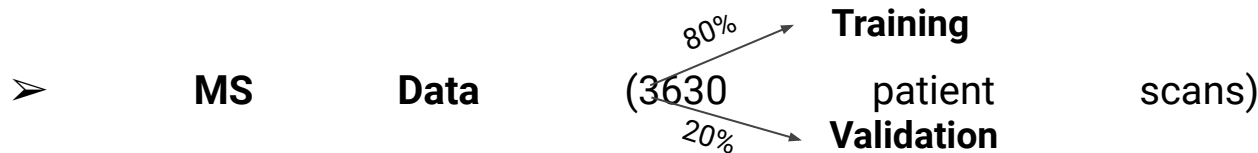


- Weighted binary cross entropy was used as loss function.

Experimentation (First Phase: Pre-training)



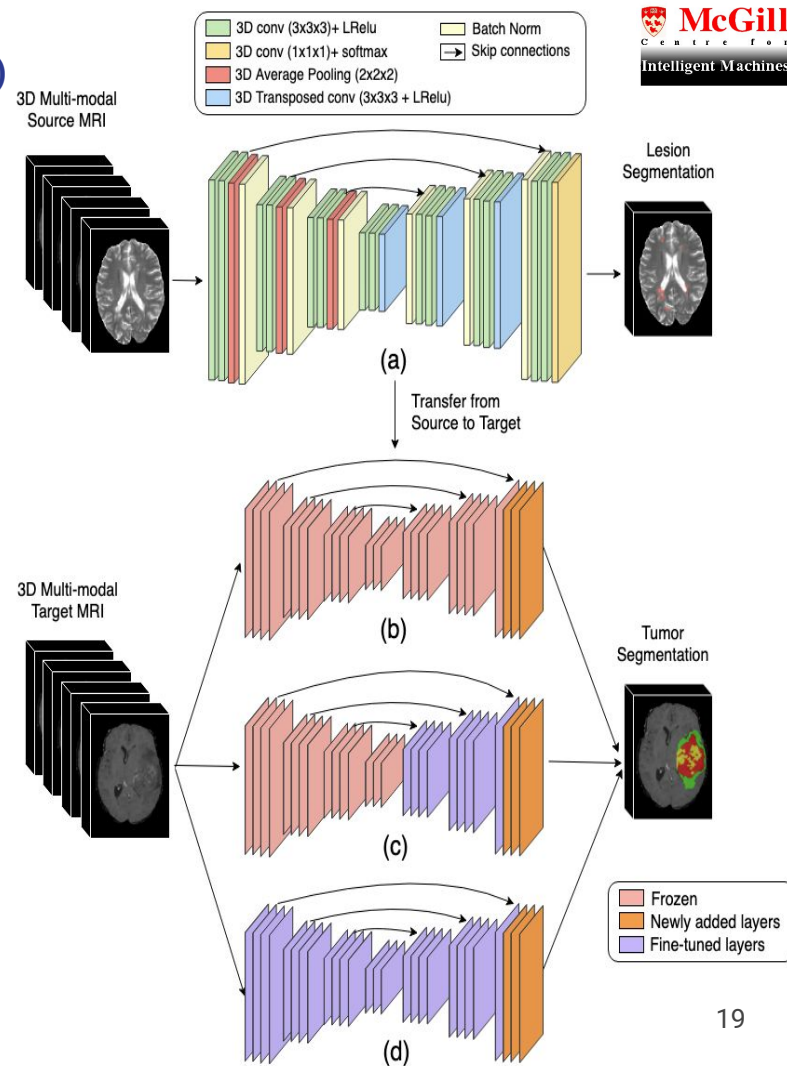
Pre-training the UNet with source MS data for T2 lesion segmentation.



- 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:

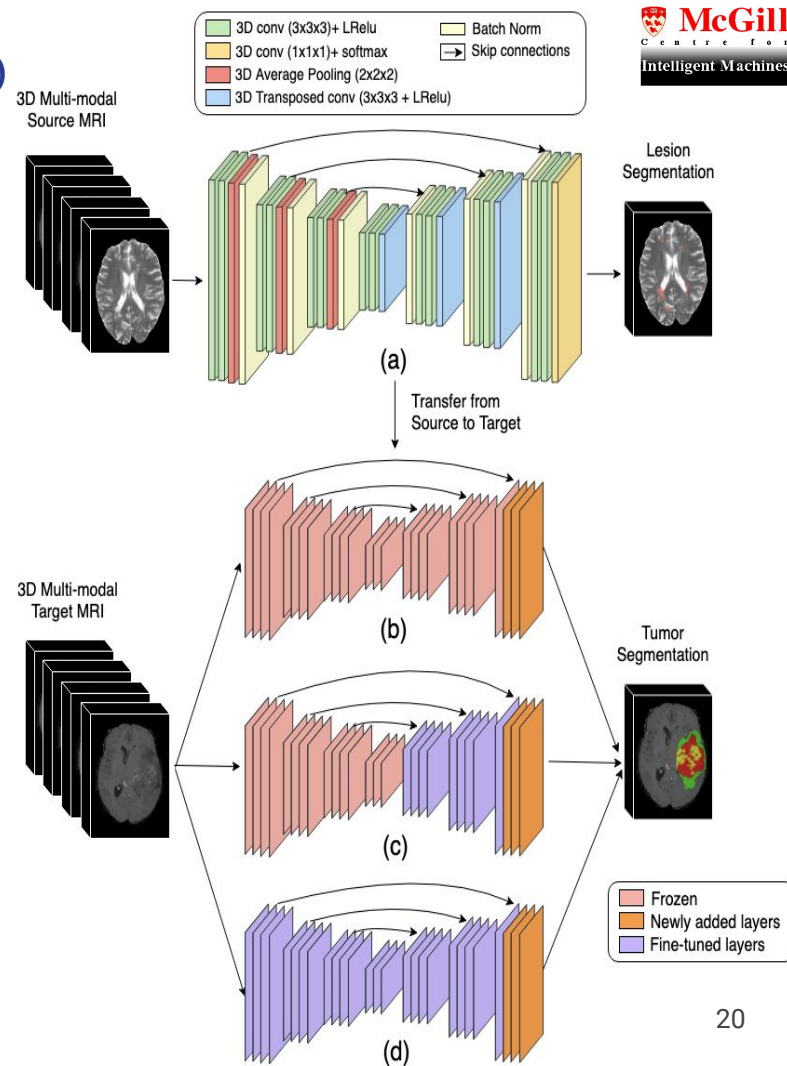


Experimentation (Second Phase: Fine-tuning)

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

★ Transfer Learning

- FT_Last Three
- FT_Decoder
- FT_All



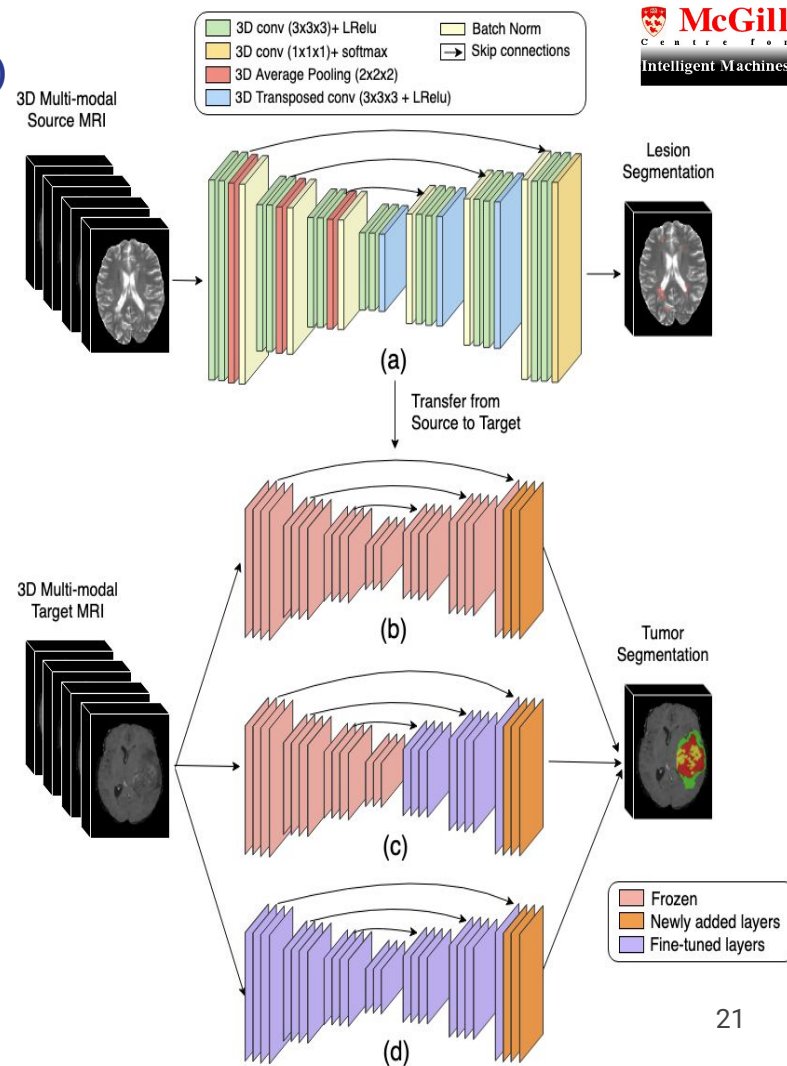
Experimentation (Second Phase: Fine-tuning)

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)

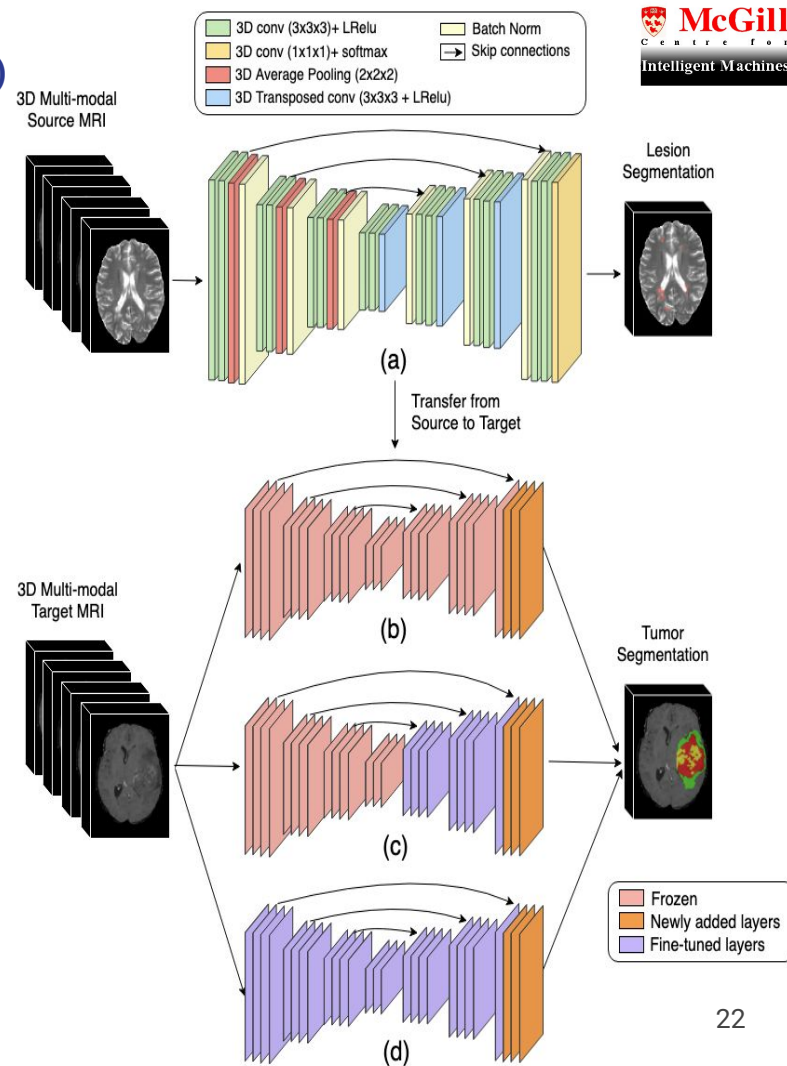
For 20, 50, 100, 150 brain tumor MRI scans:

★ Transfer Learning

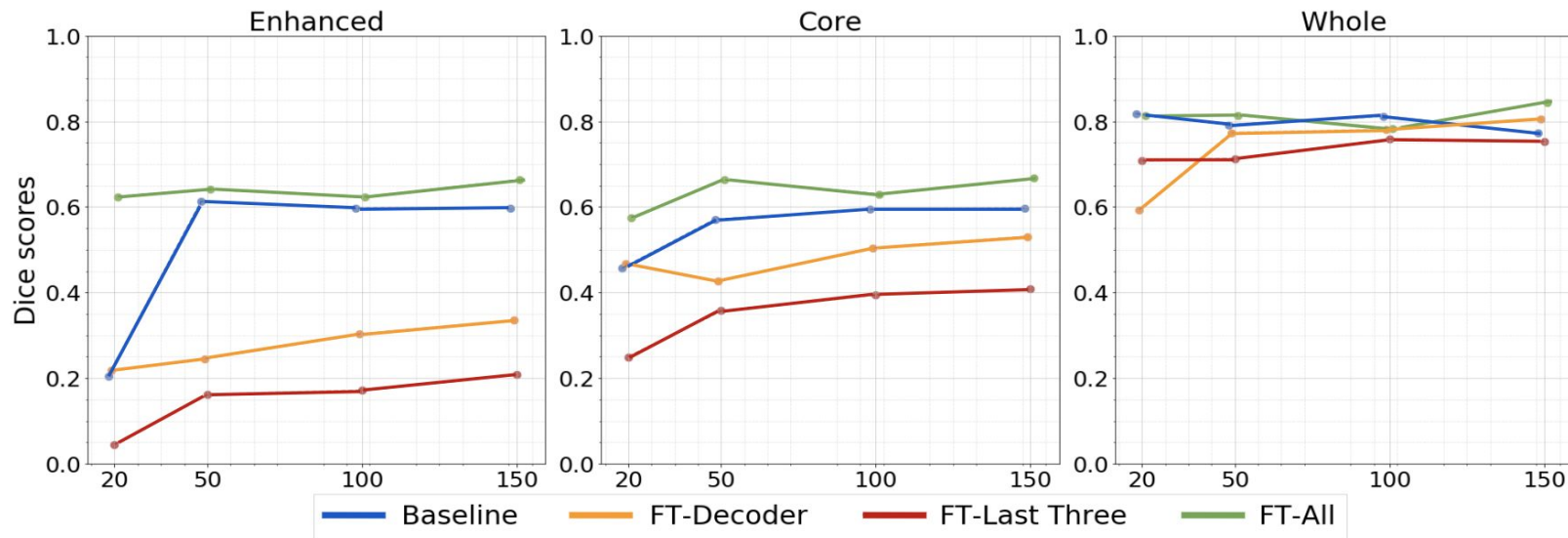
- FT_Last Three
- FT_Decoder
- FT_All

★ 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.

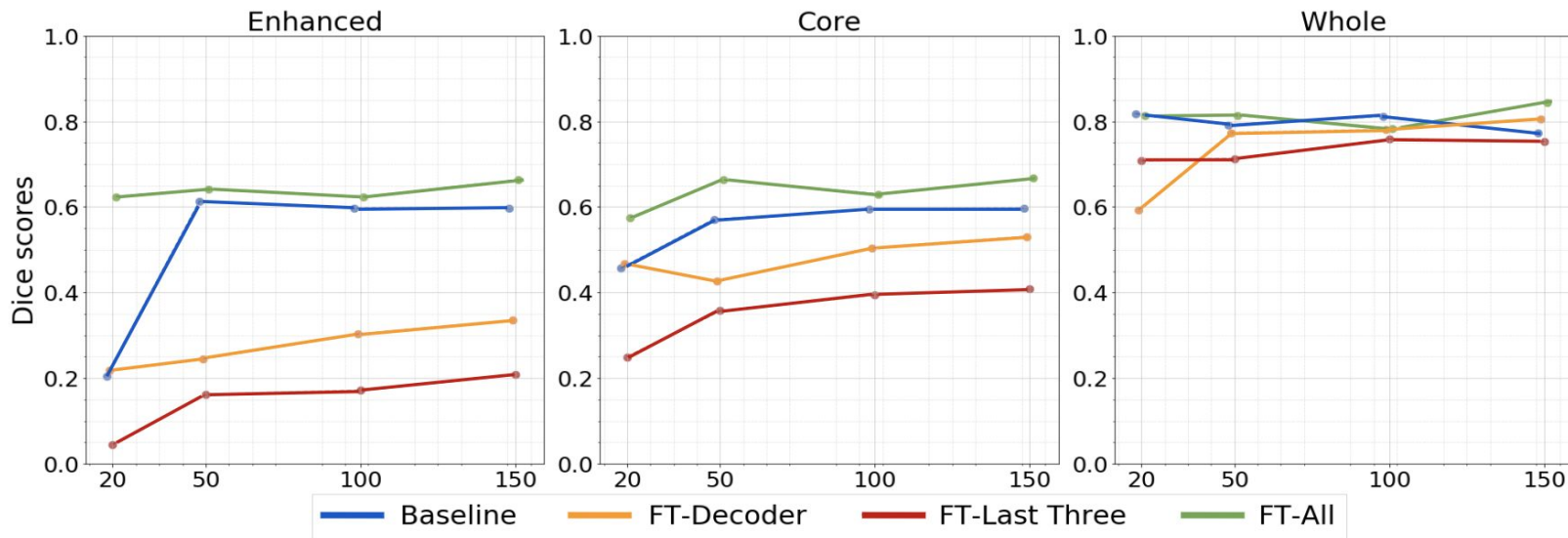


Quantitative Results (on BraTS 2018 Validation set)



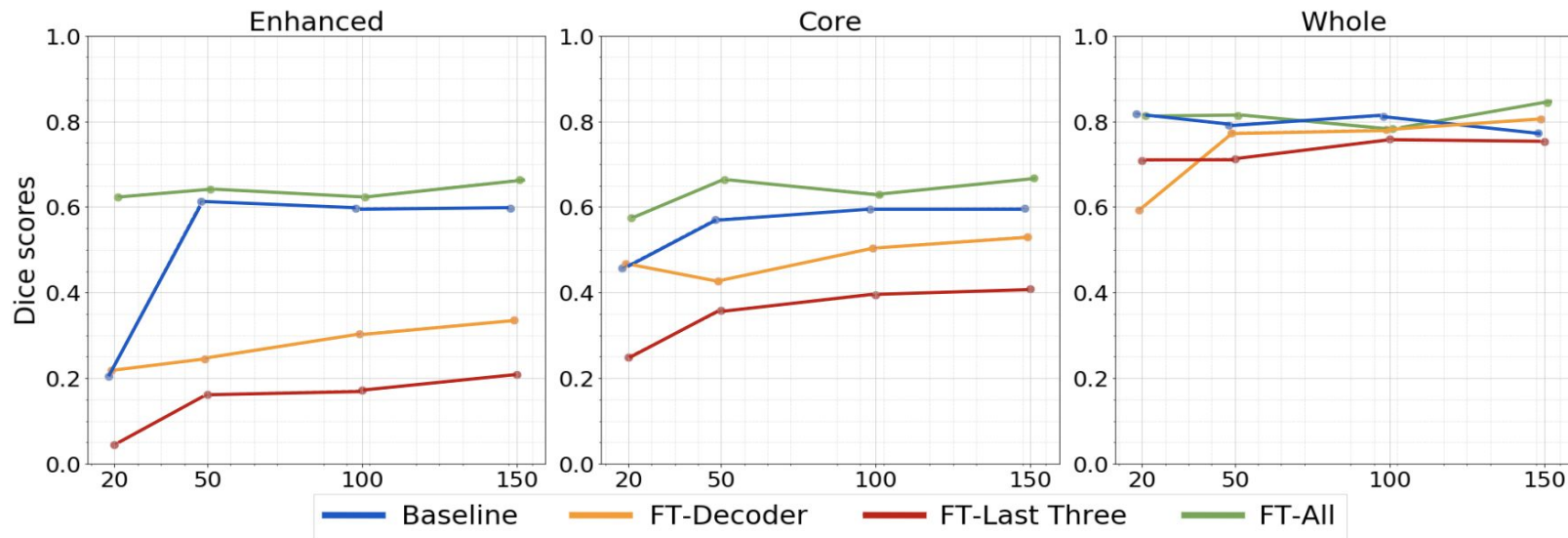
➤ **FT-All outperforms the baseline** in almost every case.

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)

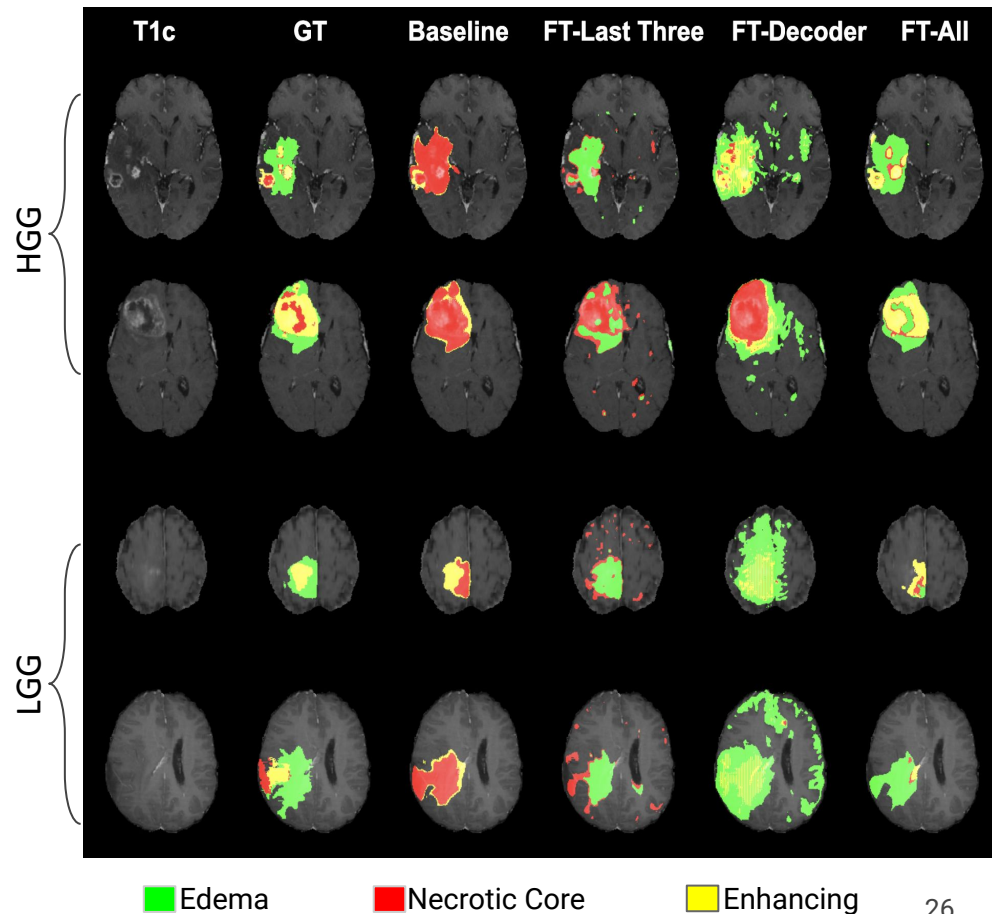


- **FT-All outperforms the baseline** in almost every case,
- Best when the **number of tumor cases is extremely low**, i.e. 20.
- As the number of brain tumor samples increase, **the gain of FT-All over baseline diminishes**.

Qualitative Results

Fine-tuned with 20 brain tumor cases

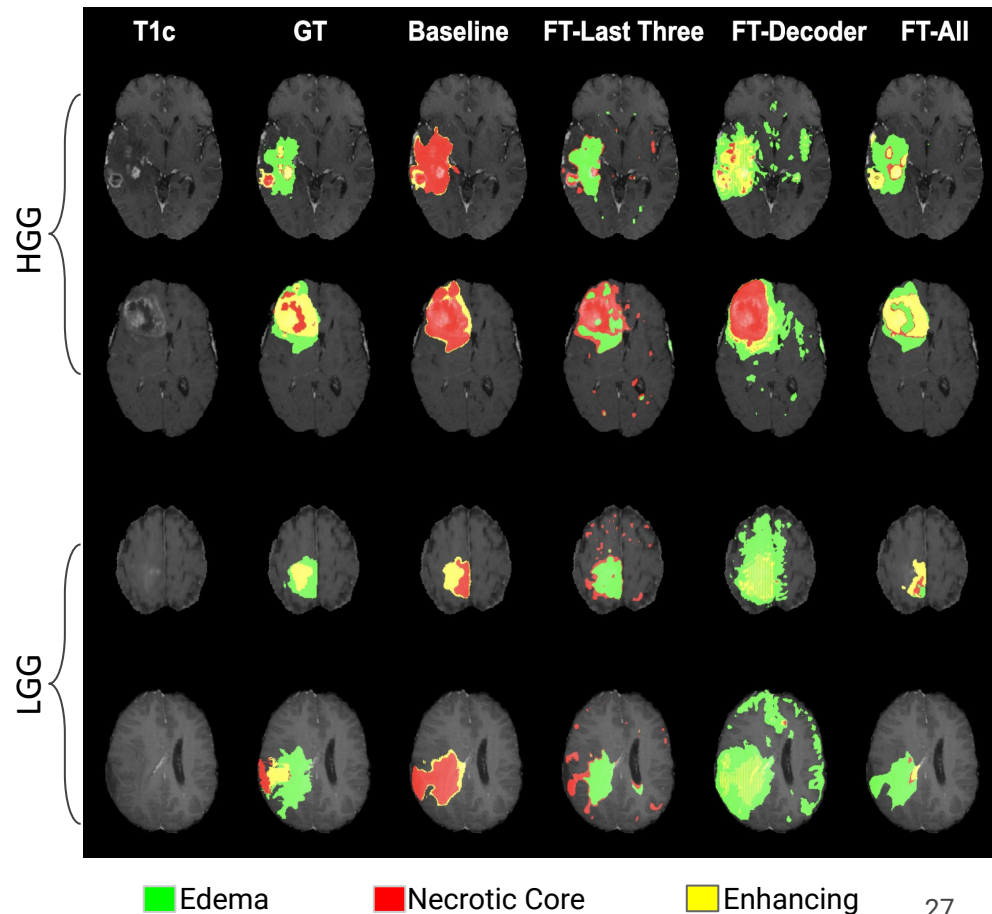
- **FT-All is able to capture sub-structures of tumor better than the other methods.**



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^[2]
- ❖ N3 bias field inhomogeneity correction^[3]
- ❖ Nyul image intensity normalization^[4]
- ❖ 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 ^[1]

- ❖ Skull stripping
- ❖ Co-registration
- ❖ Registration to same space as source data using ANTs tool^[5]
- ❖ 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