# Segmentation of pathologies in Human Brain MRI's with uncertainty

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### Segmentation In Medical Imaging with Deep Learning (1/2)

Segmentation of brain MRIs on various datasets using Deep Learning algorithm

Quantifying the uncertainty of segmentations provided by the neural network leads to a better interpretation by medical teams

Quantifying uncertainty in deep learning remains a key unresolved issue

Implementation of several methods including Deep Ensemble and Monte Carlo Dropout





### Segmentation In Medical Imaging with Deep Learning (2/2)

- State of the Art and Uncertainty Metrics
- ❖ iSeg-2017 : 6-month infant brain MRI Segmentation
- **\*** Experimentations
- Achievements This Semester
- Future Work



### Quantifying Prediction Uncertainty (1/2)

#### **Estimating Prediction Uncertainty:**

#### **Average Probability Image:**

- Mean probability for each pixel across all model predictions
- > Central estimate of the segmentation
- ightharpoonup Formula :  $\overline{X}_i = \frac{1}{N} \sum_{j=1}^N X_{ij}$

#### **Standard Deviation Map:**

- Standard deviation for each pixel across all model predictions
- > Highlights variability and uncertainty in predictions
- > Formula:

$$\sigma_i = \sqrt{rac{1}{N}\sum_{j=1}^N (X_{ij} - \overline{X}_i)^2}$$

#### Steps:

- Mean probability calculation for each pixel from all models
- Standard deviation calculation for each pixel from all models

Provides comprehensive view of prediction reliability.

#### Formulas:

**♦ Mean Probability** for each pixel :

$$\overline{X}_i = rac{1}{N} \sum_{j=1}^N X_{ij}$$

**Standard Deviation:** 

$$\sigma_i = \sqrt{rac{1}{N}\sum_{j=1}^N (X_{ij} - \overline{X}_i)^2}$$



### Quantifying Prediction Uncertainty (2/2)

#### **Estimating Prediction Uncertainty with Shannon Entropy**

- Entropy calculated for each pixel across all model predictions
- Measures the unpredictability and information content of the segmentation
- � Formula :  $H(X_i) = -\sum_{c=1}^C p_{ic} \log_2(p_{ic})$

#### Where:

- $X_i$  is the pixel i
- ullet  $p_{ic}$  is the predicted probability for class c at pixel i
- *C* is the total number of classes

#### Steps:

- Calculate the probability distribution p for each class c at each pixel
- Compute the Shannon entropy for each pixel using the probability distribution

Provides a detailed view of prediction uncertainty based on the distribution of predicted probabilities



### State of the Art and Uncertainty Metrics: Monte Carlo Dropout

### **Monte Carlo Dropout Technique**

❖ Developed by: Yarin Gal and Zoubin Ghahramani (2016)

#### **Overview:**

- ❖ **Dropout Regularization**: Randomly deactivate neurons during each forward pass to prevent overfitting
- ❖ **Bayesian Approximation**: Treats each forward pass as a sample from a Bayesian posterior distribution by applying dropout during both training and testing
- ❖ **Predictive Distribution**: Perform multiple forward passes (30-100) for each input to generate a distribution of predictions



### State of the Art and Uncertainty Metrics: Monte Carlo Dropout

### **Key Steps**:

- **\*** Training:
  - > Train with dropout enabled (e.g., 0.4 dropout rate).
- **\*** Testing/Inference :
  - ➤ Keep dropout enabled and perform multiple forward passes (e.g., 100).
- **Aggregation**:
  - Calculate mean and variance of predictions



### State of the Art and Uncertainty Metrics: Deep Ensembles

#### **Deep Ensembles Technique**

**♦ Developed by**: Balaji Lakshminarayanan, Alexander Pritzel, and Charles Blundell (2017)

#### Overview:

- Multiple Models: Train multiple independent neural networks with different initializations
- **Ensemble Predictions**: Each model makes a separate prediction for the same input
- Robustness: Aggregating predictions from different models enhances robustness and captures uncertainty

### **Key Steps:**

- Training:
  - Train 5 networks separately with different initializations
- **Prediction**:
  - Each network makes its own prediction
- **♦** Aggregation :
  - Compute mean and variance of predictions



### Construction of the Final Prediction in Binary Segmentation

#### **Model Output:**

- **Probabilities**: Each pixel is assigned a probability *p* (between 0 and 1) indicating the confidence that the pixel belongs to the target class
- Shape: For an input image of dimensions (H, W) the model output is also (H, W), with each value representing a probability

#### **Classification Threshold:**

- **Threshold** t: Probabilities are converted into binary classification using a threshold, typically t = 0.5
- **Decision**:
  - ► If  $p \ge t$ : the pixel is classified as the target class
  - ightharpoonup If p < t: the pixel is classified as the background class

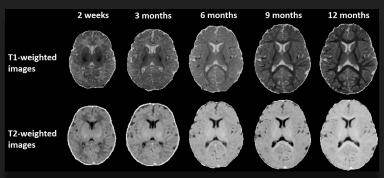
#### **Result Interpretation:**

- ❖ **Probabilities**: Display the predicted probabilities for each pixel
- ♦ Binary Predictions : Binary classification of each pixel using the threshold t. (1 = target class, 0 = background class)



### iSeg-2017: 6-month infant brain MRI Segmentation

- iSeg-2017 challenge focuses on comparing semi-automatic algorithms for segmenting 6-month infant brain MRIs using T1 and T2 images
- ☐ Critical for studying the dynamic first year of postnatal human brain development and associated cognitive and motor functions
- Intense phase at 6 months presents the lowest tissue contrast, posing significant challenges for accurate segmentation
- Engages researchers to develop and test automatic segmentation algorithms for white matter, gray matter, and cerebrospinal fluid



**Figure 1 :** MIICCAI Grand Challenge on iSeg-2017, 6-Month infant Brain MRI Segmentation, <u>iSeg-2017</u>



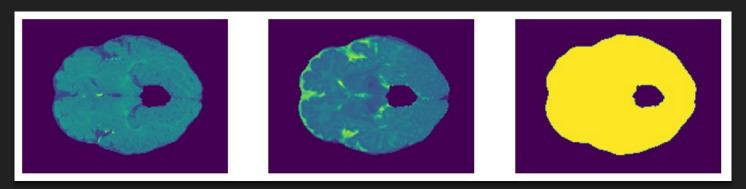


Figure 2: Input for the segmentation of the brain area

#### **Deep Ensembles for Uncertainty Estimation**

- ❖ 5 networks are trained separately
- **♦** Each network is independently initialized

#### **Monte Carlo Dropout for Uncertainty Estimation**

❖ 100 predictions with Dropout Rate of 0.4



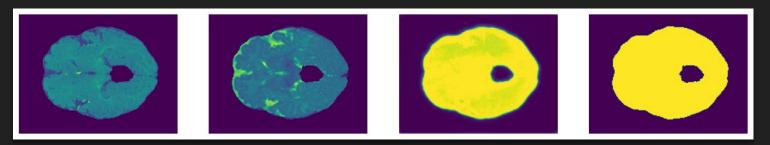


Figure 3: Mean prediction with Deep Ensembles Method, for Patient 1, slice sz // 2

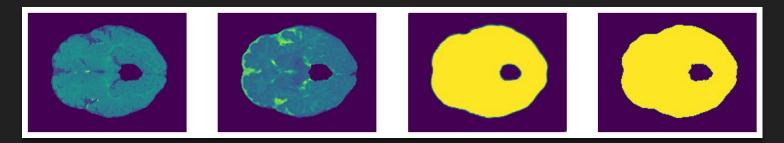


Figure 4: Mean prediction with Monte Carlo Dropout Method for Patient 1, slice sz // 2



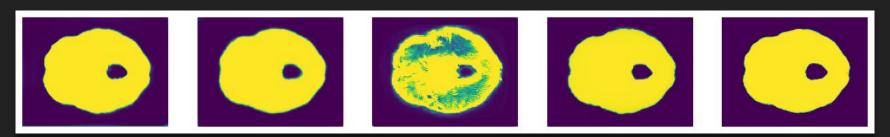


Figure 5: Predictions from the 5 Networks in the Ensemble

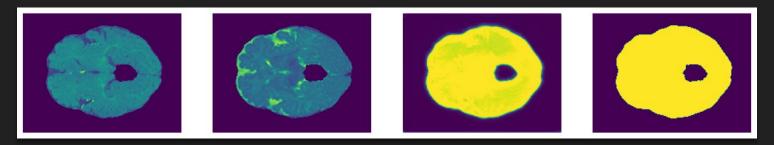
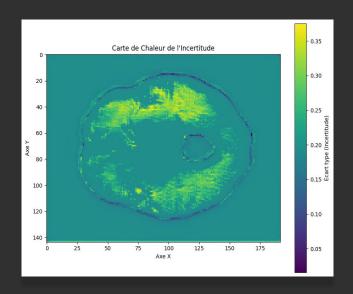


Figure 6: Mean prediction with Deep Ensembles Method, for Patient 1, slice sz // 2



### **Experimentations: Segmentation of Brain Area with Deep Ensembles**



**Figure 7 :** Standard Deviation Map for Deep Ensembles

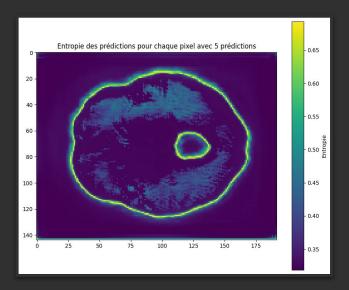
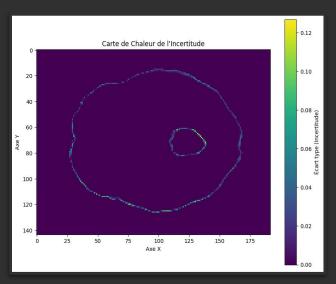


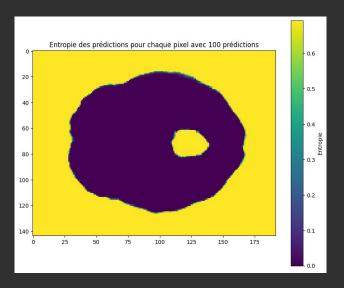
Figure 8 : Shannon Entropy for Deep Ensembles



### **Experimentations: Segmentation of Brain Area with Monte Carlo Dropout**



**Figure 9 :** Standard Deviation Map for Monte Carlo Dropout



**Figure 10 :** Shannon Entropy for Monte Carlo Dropout

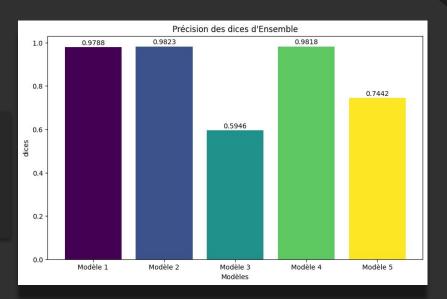


#### Dice Formula:

$$D = rac{2 imes |X \cap Y|}{|X| + |Y|}$$

where:

- $|X\cap Y|$  represents the number of elements common to both sets X and Y.
- |X| and |Y| are the respective sizes of sets X and Y.



**Figure 11 :** Dice coefficients of the 5 networks for the same input



### Experimentations: Segmentation of White and Gray Matter

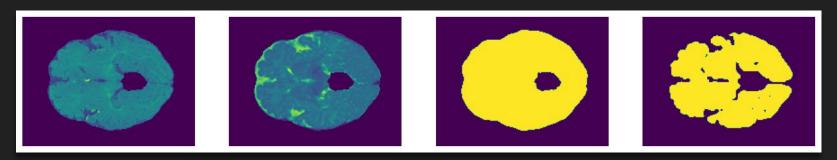


Figure 12: Input for the segmentation of White and Gray matter

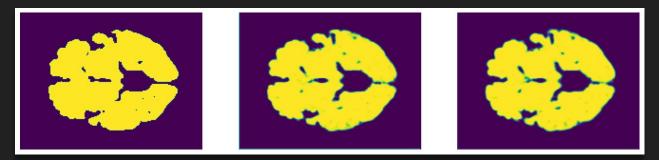
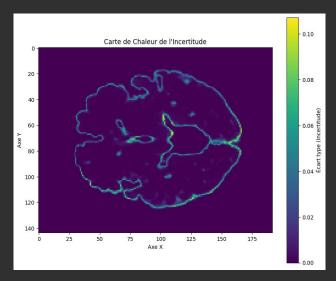


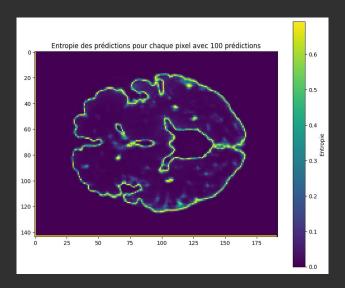
Figure 13: GroundTruth vs Mean Prediction for Deep Ensembles vs Monte Carlo Dropout



### Experimentations: Segmentation of White and Gray Matter with Monte Carlo Dropout



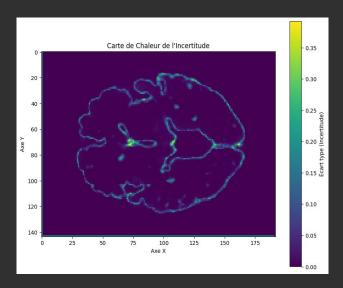
**Figure 14 :** Standard Deviation Map for Monte Carlo Dropout



**Figure 15 :** Shannon Entropy for Monte Carlo Dropout



### Experimentations: Segmentation of White and Gray Matter with Deep Ensembles



**Figure 16 :** Standard Deviation Map for Deep Ensembles

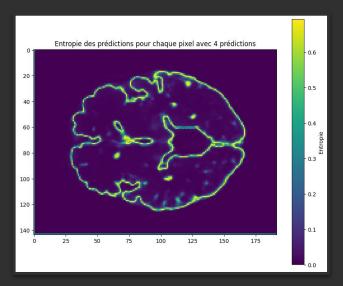
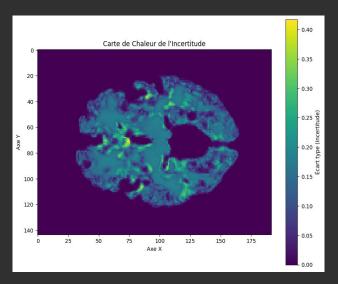


Figure 17: Shannon Entropy with Deep Ensembles



### Experimentations: Segmentation of White Matter with Deep Ensembles



**Figure 18 :** Standard Deviation Map for Deep Ensembles

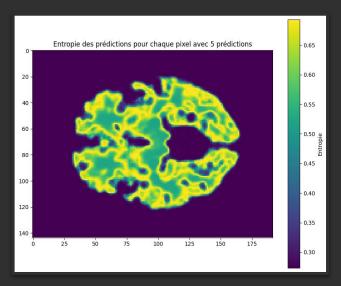
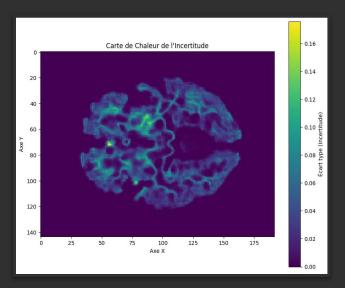


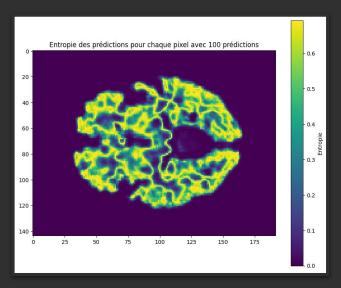
Figure 19: Shannon Entropy with Deep Ensembles



### Experimentations: Segmentation of White Matter with Monte Carlo Dropout



**Figure 20 :** Standard Deviation Map with Monte Carlo Dropout



**Figure 21 :** Shannon Entropy with Monte Carlo Dropout



### Related Work: Achievements for this Semester

#### **Experimentation**:

- Experimented with two methods for uncertainty quantification
- Compared the effectiveness of both methods

#### **Segmentation**:

Performed basic segmentation tasks using a U-net model

### **Uncertainty Measures**:

Implemented basic uncertainty measures including mean, standard deviation, and Shannon entropy



### Future Work: Research Focus for Next Semester

#### **Identifying Sources of Uncertainty:**

- Differentiating between aleatoric uncertainty (data-related) and epistemic uncertainty (model-related)
- ❖ Objective: Determine which type of uncertainty is being measured and identify its source

#### Dataset:

Work with MRBrains dataset

#### **Complex Segmentations**:

❖ Perform segmentations on even more complex structures



### References

- [1]: *Mathématiques et imagerie*, Bibliothèque Tangente n°77, Edition Pôle, 2022 <a href="https://infinimath.com/librairie/pdf/BIB77\_sommaire.pdf">https://infinimath.com/librairie/pdf/BIB77\_sommaire.pdf</a>
- [2]: *Brain Tumor Segmentation and Survival Prediction using 3D Attention UNet*, Mobarakol Islam, Vibashan, Jeya Maria Jose, Navodini Wijethilake, Uppal Utkarsh Hongliang (PDF) Brain Tumor Segmentation and Survival Prediction Using 3D Attention UNet, Published on ResearchGate
- [3]: *Is segmentation uncertainty useful?* Steven Czolbe, Kasra Arnavaz, Oswin Krause, Aasa Feragen [2103.16265] Is segmentation uncertainty useful?, Published on Arxiv
- [4]: Simple and Scalable Predictive Uncertainty Estimation using Deep Ensembles, Balaji Lakshminarayanan Alexander Pritzel Charles Blundell [1612.01474] Simple and Scalable Predictive Uncertainty Estimation using Deep Ensembles, Published on Arxiv
- [5]: Confidence Calibration and Predictive Uncertainty Estimation for Deep Medical Image Segmentation, Alireza Mehrtash, Graduate Student Member, William M. Wells III [1911.13273] Confidence Calibration and Predictive Uncertainty Estimation for Deep Medical Image Segmentation, Published on Arxiv
- [6]: **Dropout as a Bayesian Approximation: Representing Model Uncertainty in Deep Learning,** Yarin Gal, Zoubin Ghahramani, University of Cambridge [1506.02142] Dropout as a Bayesian Approximation: Representing Model Uncertainty in Deep Learning, **Published on Arxiv**