

Unsupervised learning for

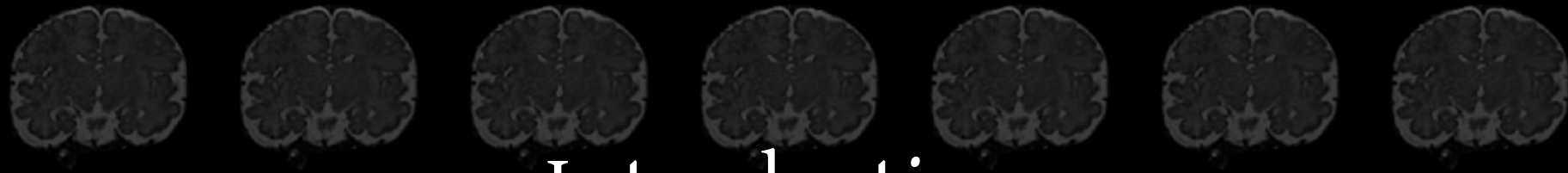
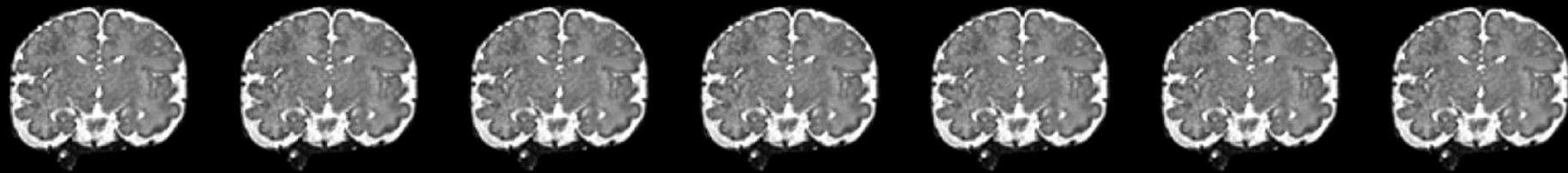
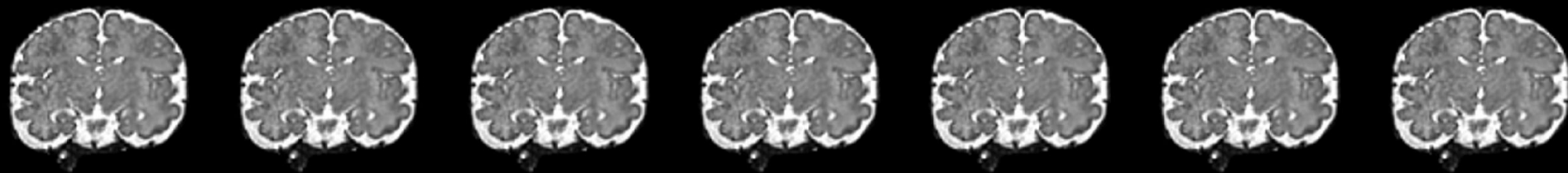
DETECTION

in fetal MRI

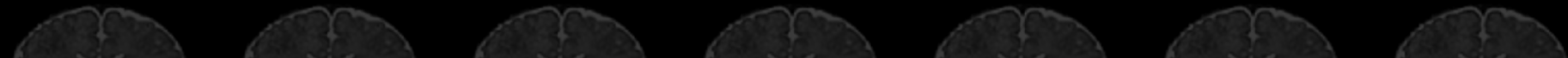


HARVARD
MEDICAL SCHOOL





Introduction



What is anomaly detection?

*“Anomaly detection (AD) aims at finding unexpected or rare events in data streams, commonly referred to as **anomalous events**”*

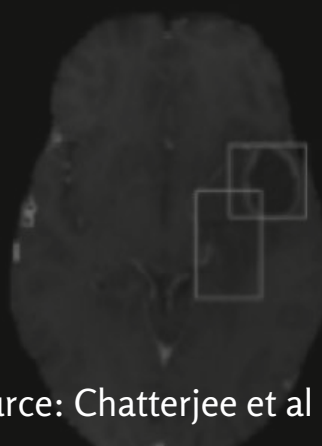
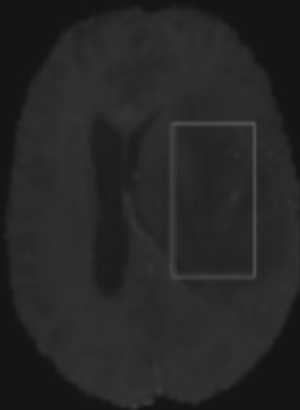
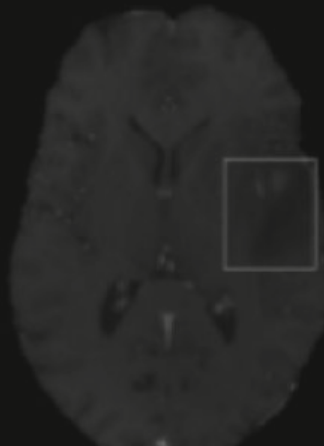
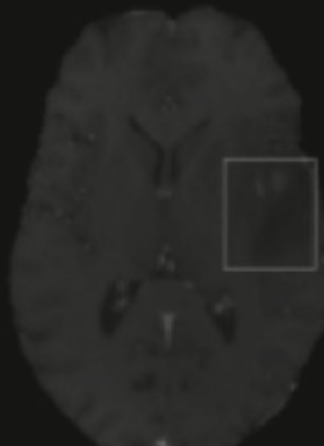
Identify relevant indicators of diseases and differentiating them from those of typical healthy tissue characteristics

Source: Schneider & Xhafa (2022)

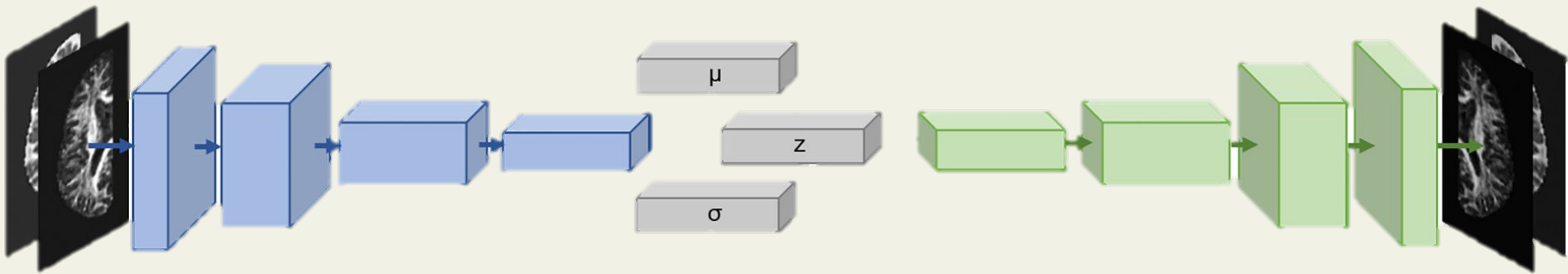
Source: Chatterjee et al (2022)

Why anomaly detection?

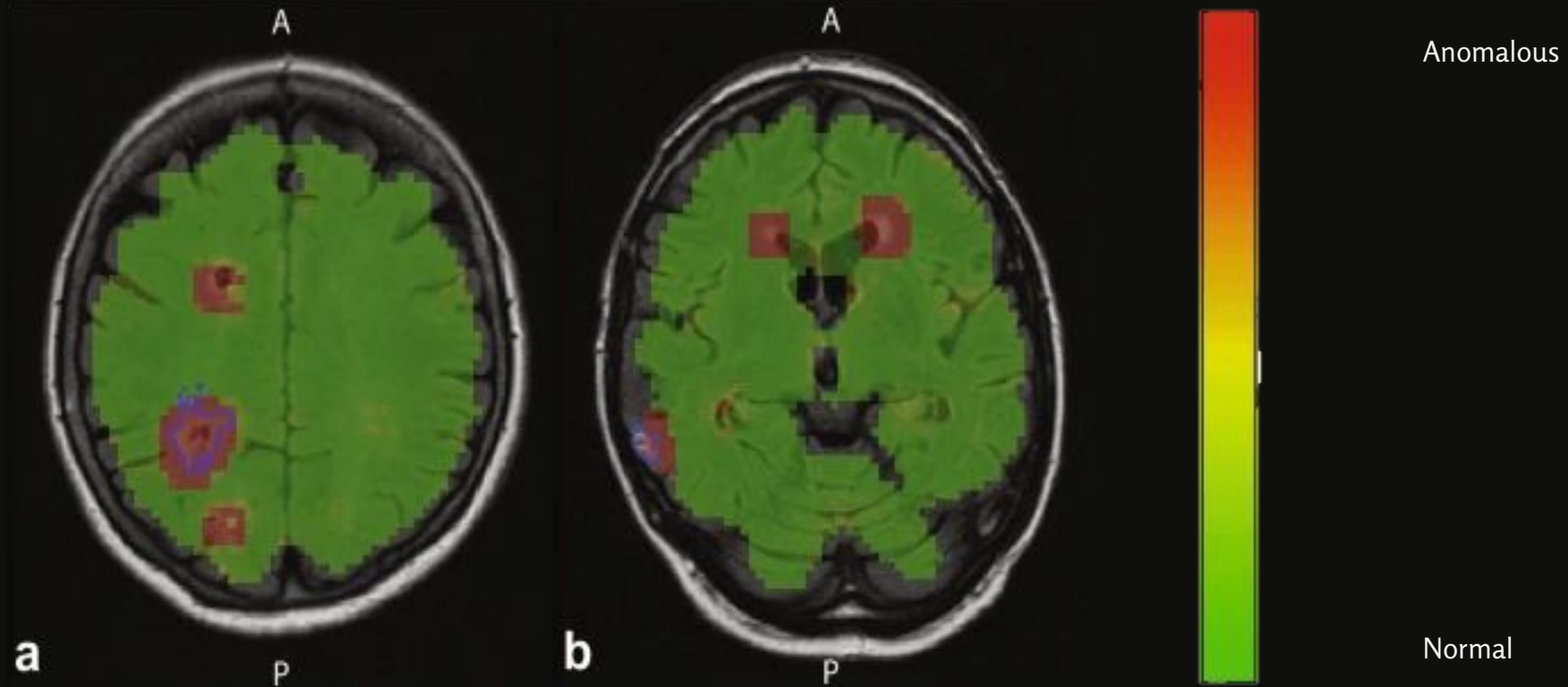
Diagnostics of brain pathology remain undiscovered in up to 5-10% of cases. Localization of anomalies in magnetic resonance imaging (MRI) can aid radiologists in pathology diagnosis.



What are unsupervised learning models?



Anomaly detection can be achieved through unsupervised learning methods.



Source: van Hespen et al (2021)

Previous work

Subtle anomaly detection: Application to brain MRI analysis of de novo Parkinsonian patients,

Muñoz-Ramirez et al (2021)

Implements Variational Auto-Encoder (VAE) model to detect brain MRI anomalies, validated with Parkinsonian patients.

StRegA: Unsupervised anomaly detection in brain MRIs using a compact context-encoding variational autoencoder

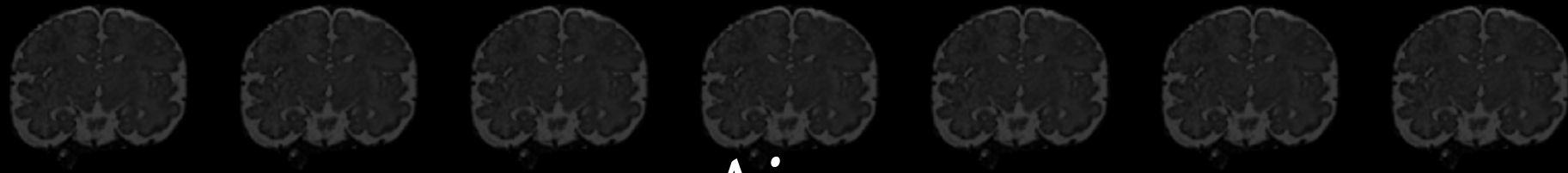
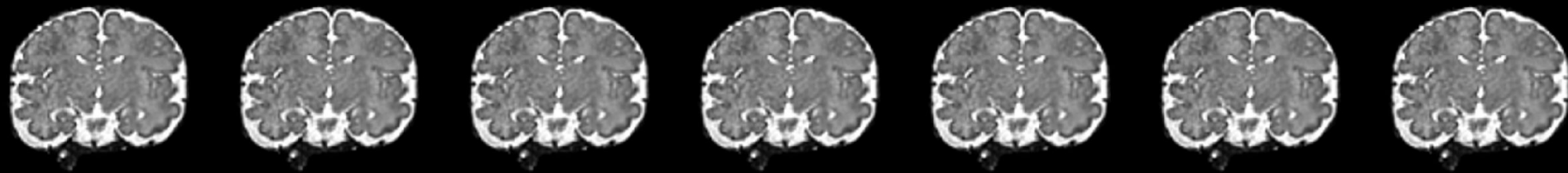
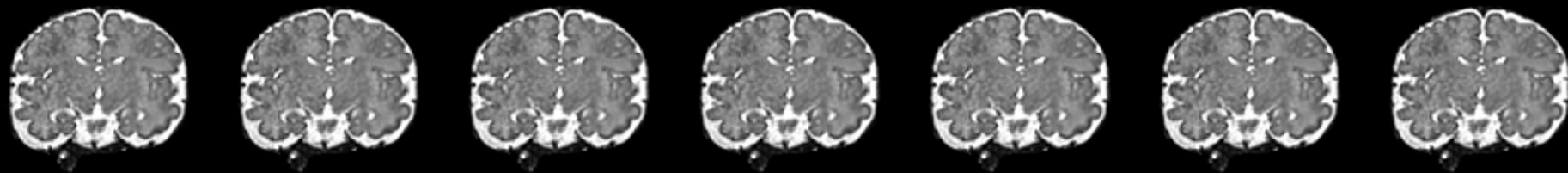
Chatterjee et al (2022)

Implements a VAE model to detect brain tumors. Adds binarization methods and morphological openings to reduce false positives.

Reversing the Abnormal: Pseudo-Healthy Generative Networks for Anomaly Detection

Bercea et al (2023)

Proposes a refinement model after a base reconstruction through an in-painting model to reduce false positives.



Aim



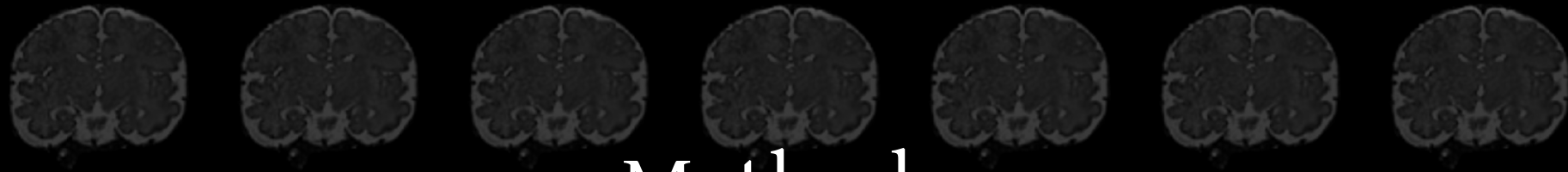
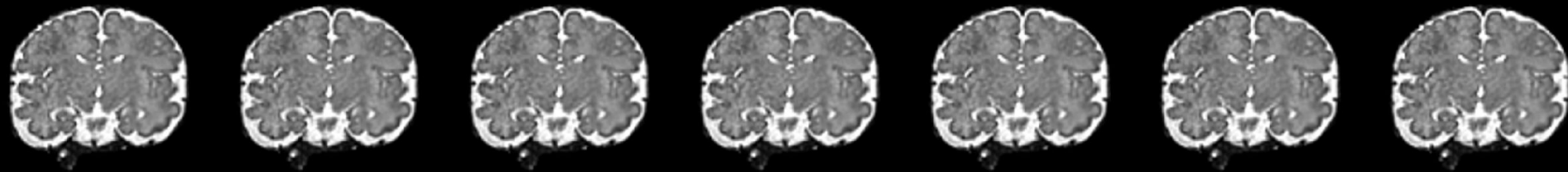
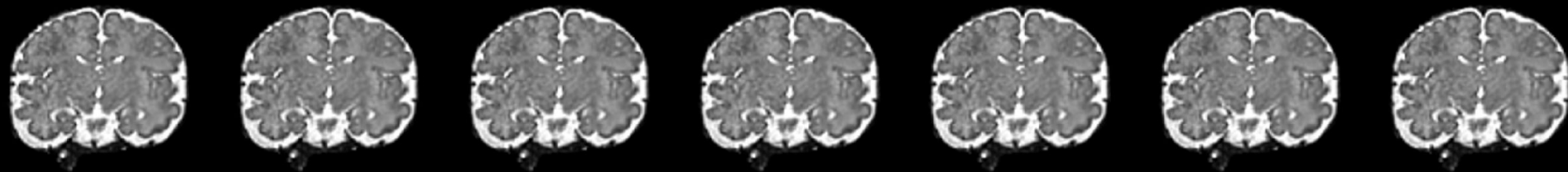
Aim

Augment accuracy of the AD process

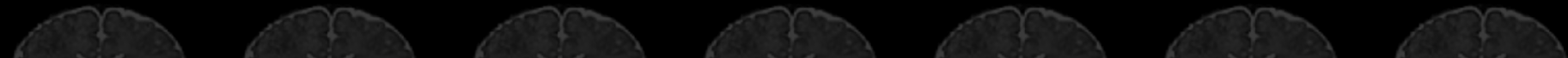
- Improve general detection performance of anomalies in fetal brain

Be able to detect not only developmental delays, but also developmental ones

- Detect true-positive anomalies by reflecting the temporal variation due to neurodevelopment during gestation



Methods



Datasets

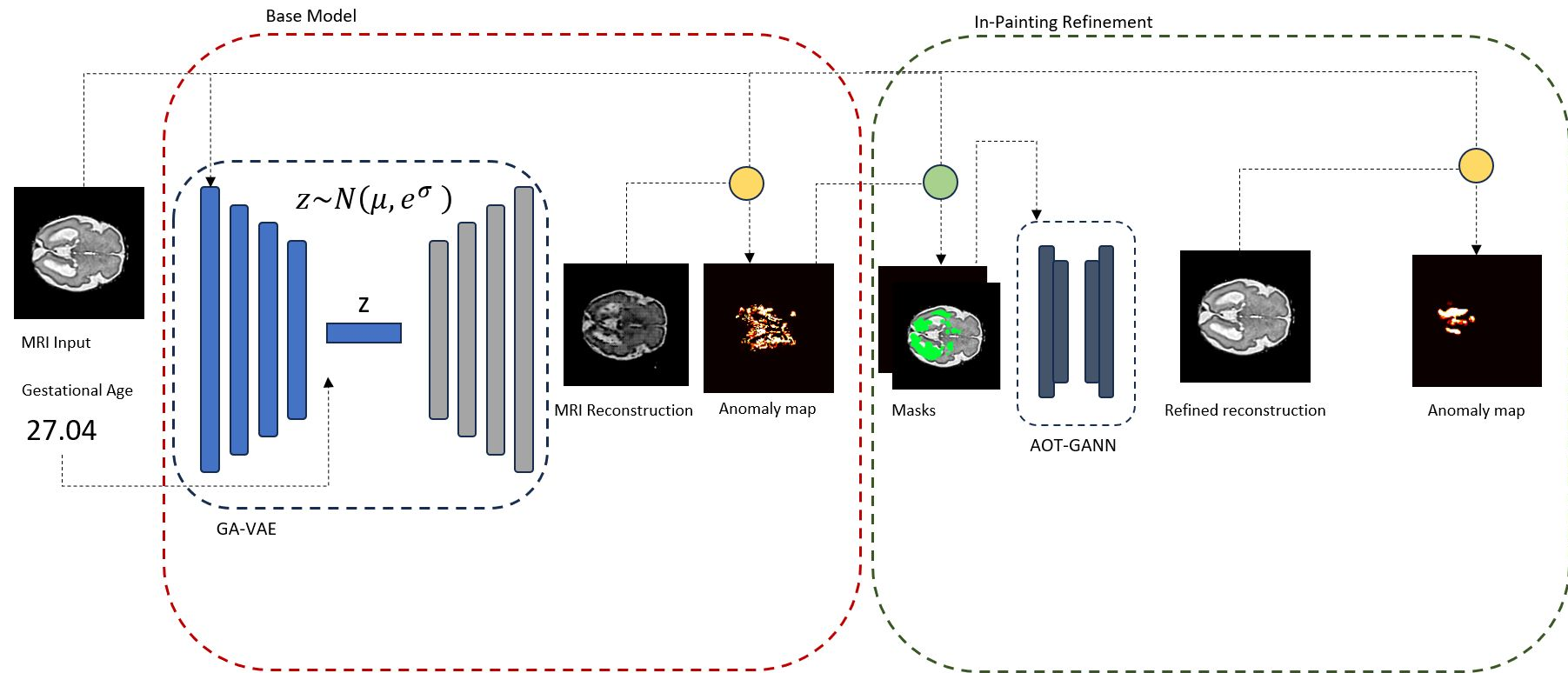
MRI data was extracted from the CHD, Placenta, TMC, VGH and BCH datasets.

Typically Developing (TD) Subjects

A total of 227 subjects. 181 used for training and 46 for testing.

Ventriculomegaly (VM) Subjects

A total of 69 subjects.



■ Conv 3x3 Stride 2, Leaky ReLU, Batch normalization

■ T. Conv 3x3 Stride 2, Leaky ReLU, Batch normalization

● $anomaly = |eq(X_{recon}) - eq(\bar{X})| \cdot loss_{per}(\bar{X}, X_{recon})$

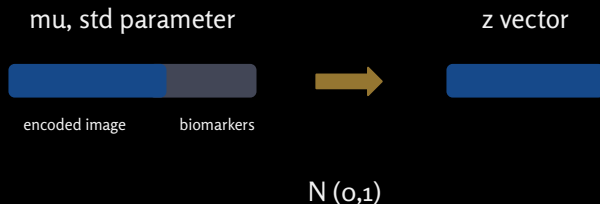
● $mask = binarize(gaussfilter(binarize(anomaly, p_{95})), 0.1)$

Providing significant novelty for fetal MRI: gestational age (GA) as input

Developmental context

Recognition of developmental delays as anomalies

Concatenation method



Refining reconstruction through in-painting model



Normal Abnormal

Only for training

Training of VAE and AOT-GAN models are done separately

VAE

- L2 loss only.

- No adversarial training

AOT-GAN

- Combination of L1, perceptual, and style losses

- Adversarial training

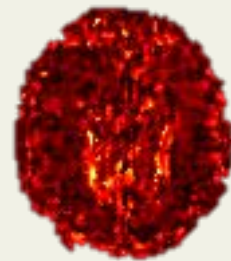
How do we define anomaly?



Input



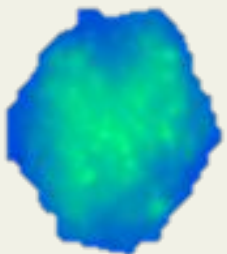
Reconstruction



Mean Squared Error
(MSE)



Mean Absolute Error
(MAE)



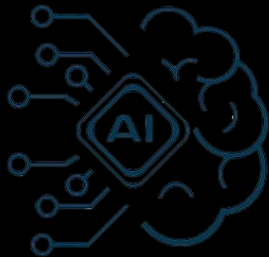
Saliency

$$anomaly = |eq(X_{rec}) - eq(\bar{X})| \cdot loss_{per}(\bar{X}, X_{rec})$$



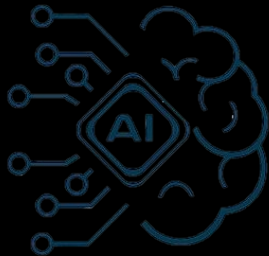
Anomaly Metric

Our current models



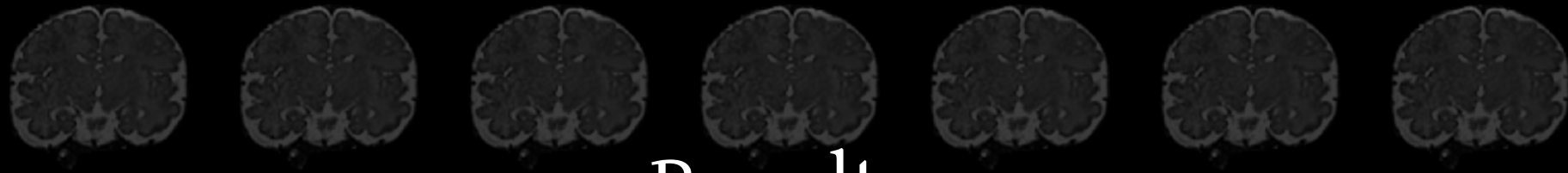
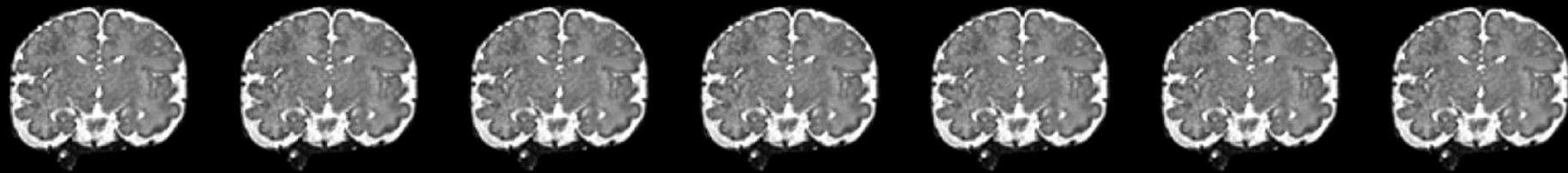
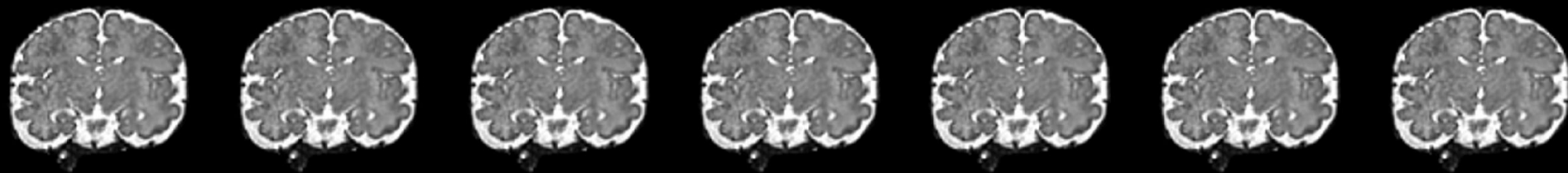
Sapi (*Quechua: Root*)

Trained for 2000 epochs. Pre-trained from a previous VAE model, it does not take GA as input. Trained with 151 subjects.

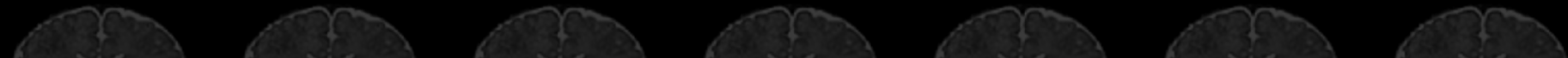


Miraywa (*Quechua: Fertility*)

Trained for 2000 epochs. Pre-trained from the Sapi model, it takes GA as input through concatenation. Trained with 227 subjects.

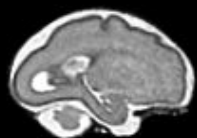


Results

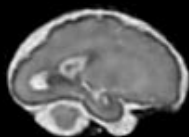


Sagittal View

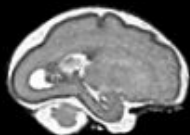
Typical Developing



Input

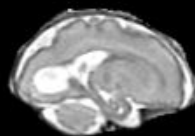


Reconstruction

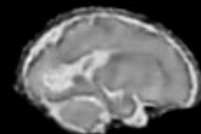


Refinement

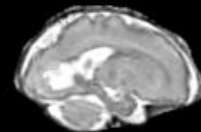
Ventriculomegaly



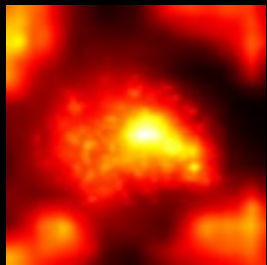
Input



Reconstruction



Refinement



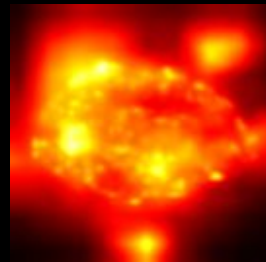
Saliency



Mask



Anomaly Metric



Saliency



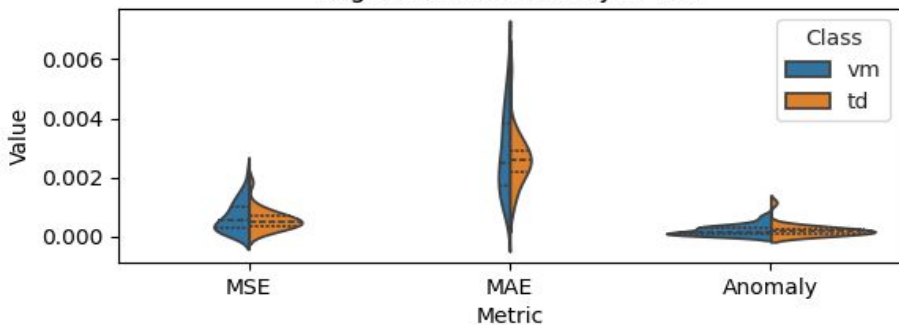
Mask



Anomaly Metric

Sapi

Sagittal Mann-Whitney U Test

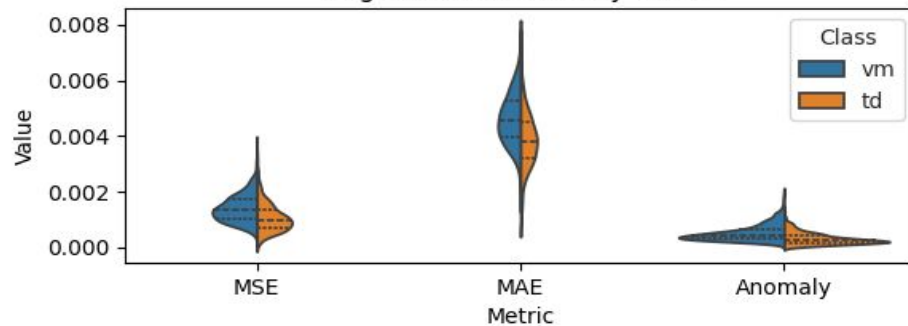


P-Values

MAE	0.8535
MSE	0.9037
Anomaly	0.8635

Miraywa

Sagittal Mann-Whitney U Test



P-Values

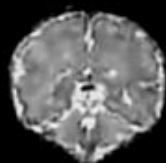
MAE	0.005
MSE	0.007
Anomaly	0.0002

Coronal View

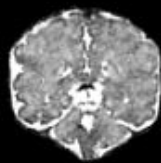
Typical Developing



Input



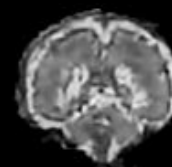
Reconstruction



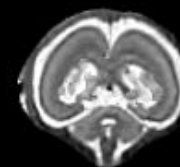
Refinement



Input

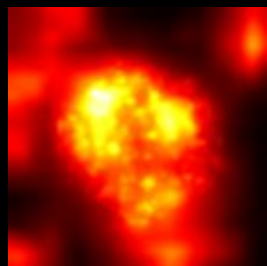


Reconstruction

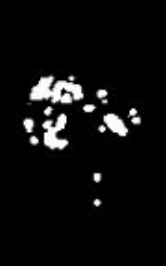


Refinement

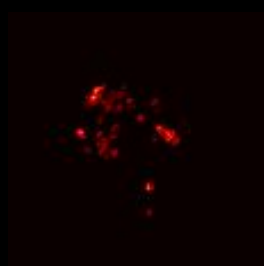
Ventriculomegaly



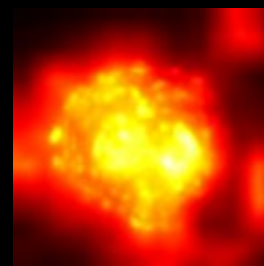
Saliency



Mask



Anomaly
Metric



Saliency



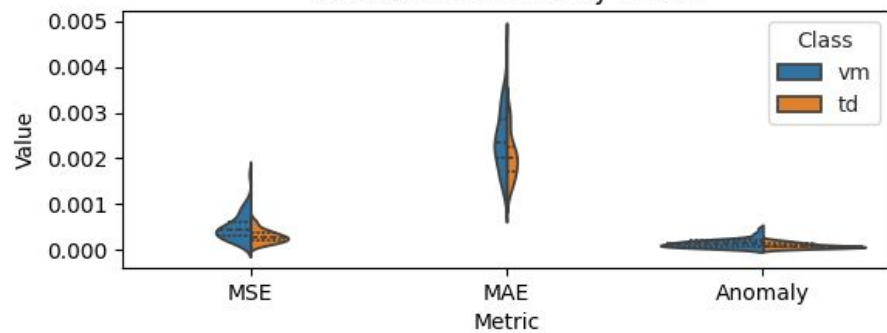
Mask



Anomaly
Metric

Sapi

Coronal Mann-Whitney U Test

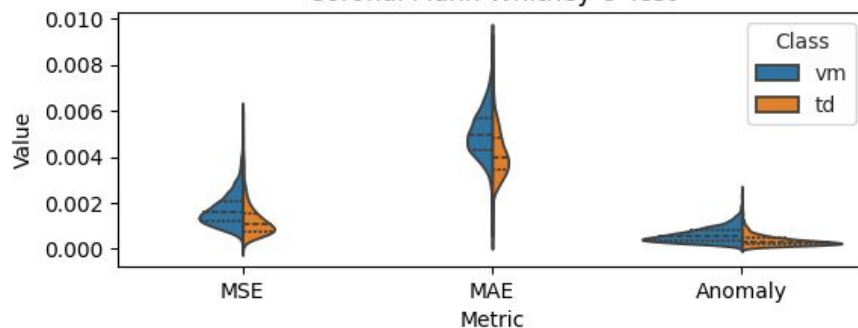


P-Values

MAE	0.001
MSE	0.01377
Anomaly	0.01329

Miraywa

Coronal Mann-Whitney U Test

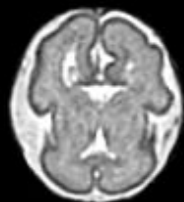


P-Values

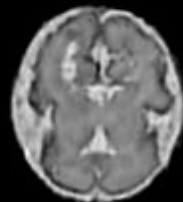
MAE	$p < 10^{-5}$
MSE	$p < 10^{-6}$
Anomaly	$p < 10^{-7}$

Axial View

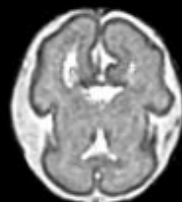
Typical Developing



Input

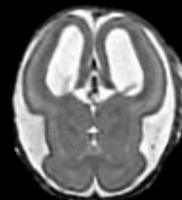


Reconstruction

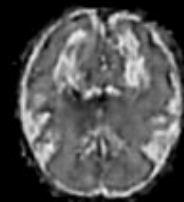


Refinement

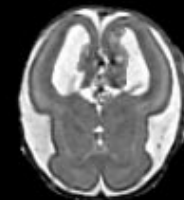
Ventriculomegaly



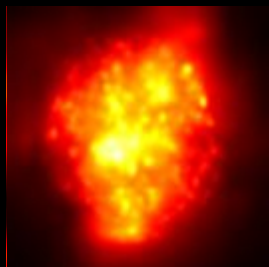
Input



Reconstruction



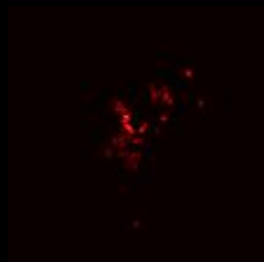
Refinement



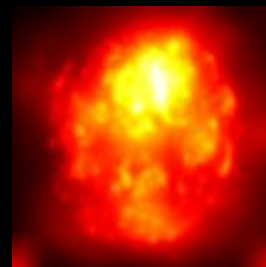
Saliency



Mask



Anomaly
Metric



Saliency



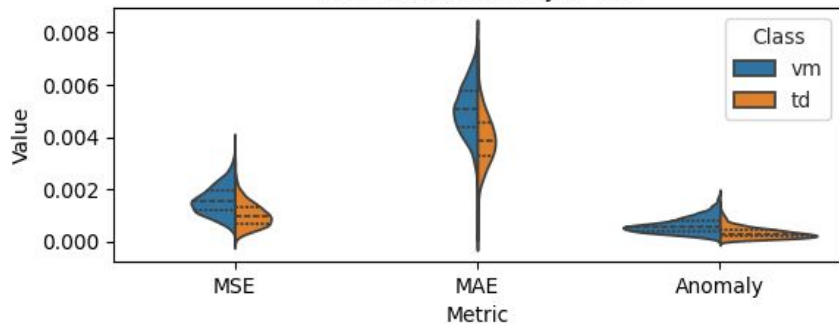
Mask



Anomaly
Metric

Sapi

Axial Mann-Whitney U Test



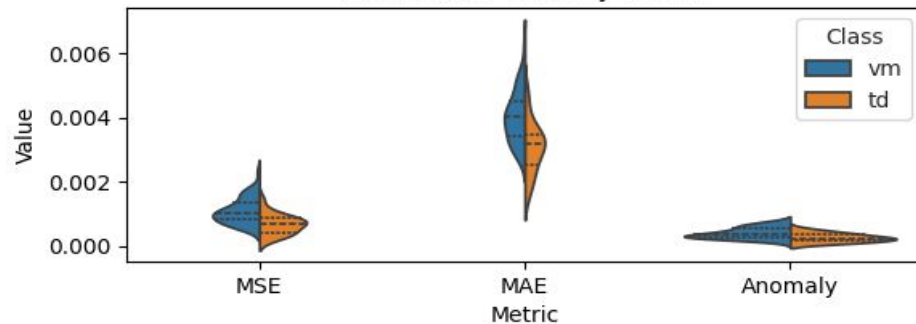
P-Values

MAE
MSE
Anomaly

$p < 10^{-5}$
 $p < 10^{-5}$
0.001

Miraywa

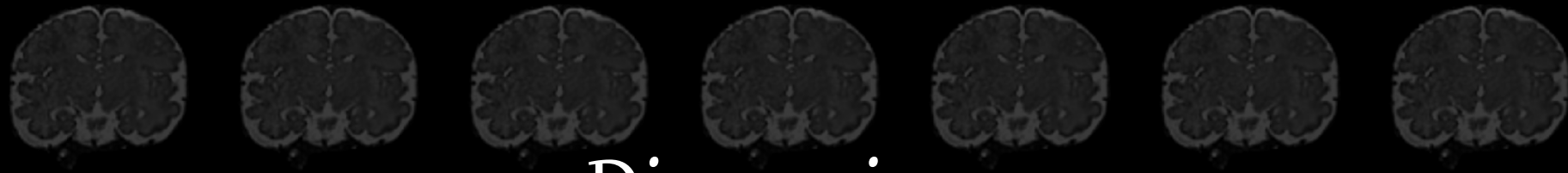
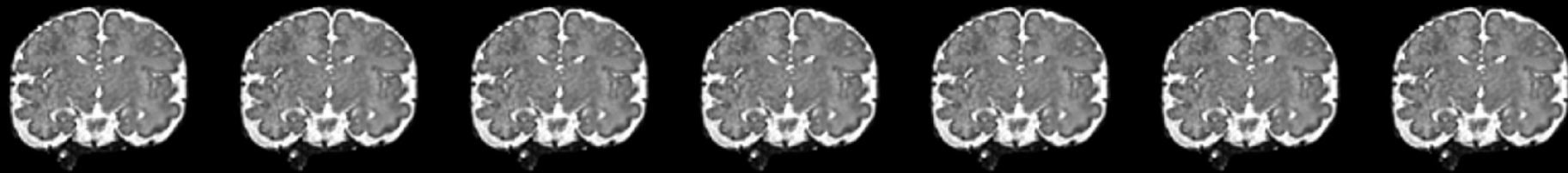
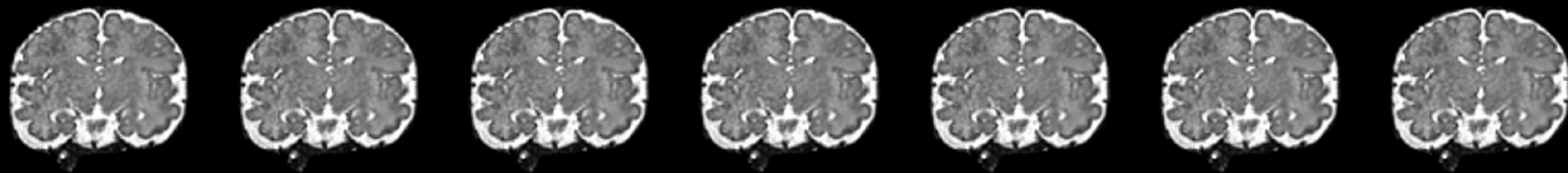
Axial Mann-Whitney U Test



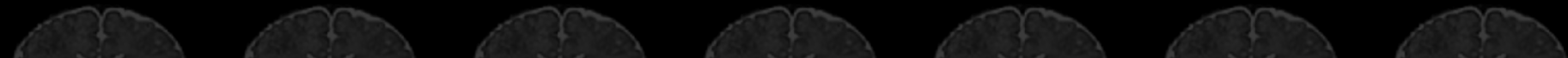
P-Values

MAE
MSE
Anomaly

$p < 10^{-7}$
 $p < 10^{-7}$
 $p < 10^{-7}$



Discussion



Limitations

Comparing models is difficult due to the difference in datasets

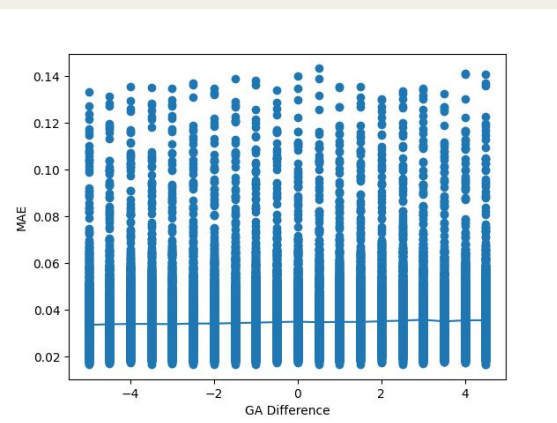
Due to the number of subject difference, it is difficult to determine if improvements are due to the dataset size or the difference in the model itself.

GA Model presents low sensitivity to GA

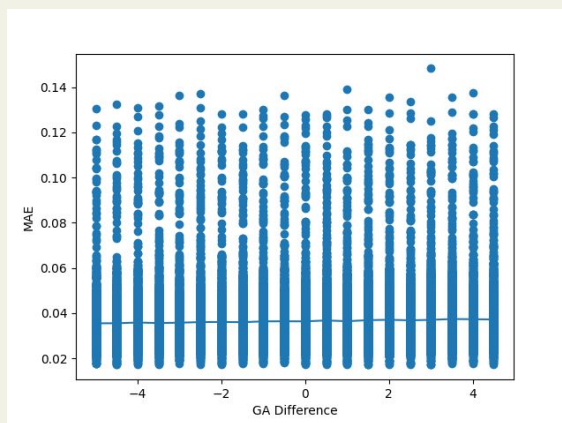
When running tests on the effects of the GA value to the errors between reconstruction and input, there is no notable influence of GA in the model.

Miraywa presented low sensitivity to GA.

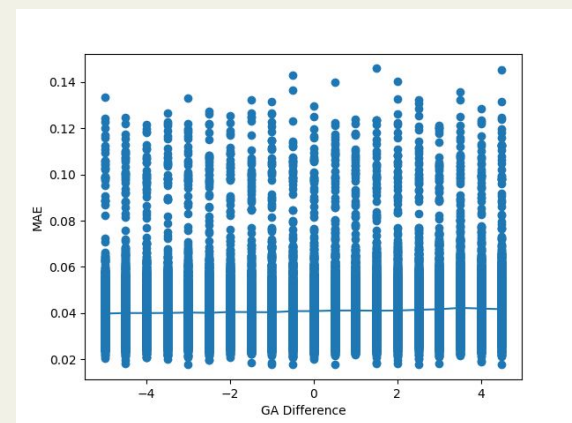
Sagittal



Coronal



Axial



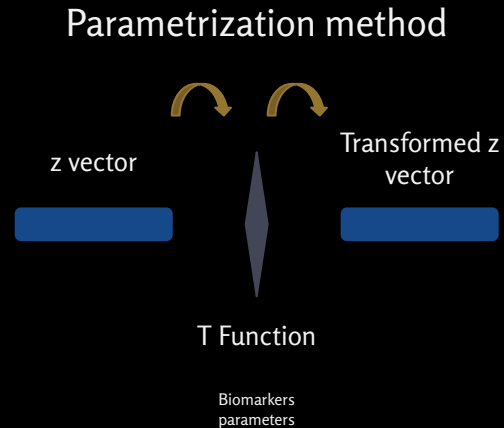
Future work

Retrain Sapi model for better model comparison

Train a non-GA model with current dataset size to properly evaluate the effect of new model

Using a new method for GA integration

Applying a parametrization method, instead of concatenation, might lead to GA holding more influence in reconstruction



Thank you.

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References

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- [2] S. Chatterjee et al., “StRegA: Unsupervised anomaly detection in brain MRIs using a compact context-encoding variational autoencoder,” *Computers in Biology and Medicine*, vol. 149. Elsevier BV, p. 106093, Oct. 2022. doi: 10.1016/j.combiomed.2022.106093.
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- [5] Bercea, C.I., Wiestler, B., Rueckert, D., Schnabel, J.A. (2023). Reversing the Abnormal: Pseudo-Healthy Generative Networks for Anomaly Detection. In: Greenspan, H., et al. *Medical Image Computing and Computer Assisted Intervention – MICCAI 2023*. MICCAI 2023. Lecture Notes in Computer Science, vol 14224. Springer, Cham. doi: 0.1007/978-3-031-43904-9_29
- [6] Y. Zeng, J. Fu, H. Chao, and B. Guo, “Aggregated Contextual Transformations for High-Resolution Image Inpainting.” *arXiv*, 2021. doi: 10.48550/ARXIV.2104.01431.