

Das Deutsche Zentrum für
Neurodegenerative Erkrankungen

Explainable Differential Diagnosis of Dementia via Contrastive Self- supervised Learning

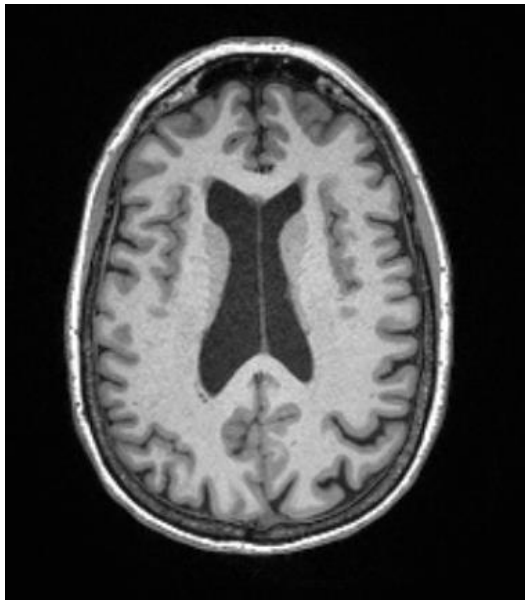
16.02.2022

Vadym Gryshchuk

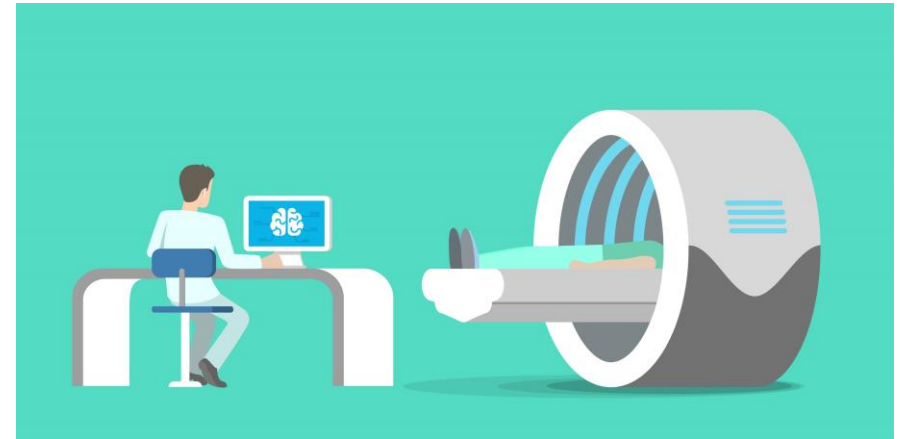


Motivation

- Early recognition of dementia
- Assistive technology for doctors



Source: <http://adni.loni.usc.edu/>, ADNI_941_S_6581



Source: <https://de.freepik.com/sentavio>, Designed by sentavio / Freepik

Outline

- Background
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- Goal
- Architecture
 - Approach
 - Evaluation
- Intermediate Results
- Conclusion & Future Steps



Outline

- Background
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- Goal
- Architecture
 - Approach
 - Evaluation
- Intermediate Results
- Conclusion & Future Steps



What is Self-supervised Learning?

- Learns **rich representations** from data without any labels/annotations



Source: <https://www.freeimages.com/photo/horse-1393073>



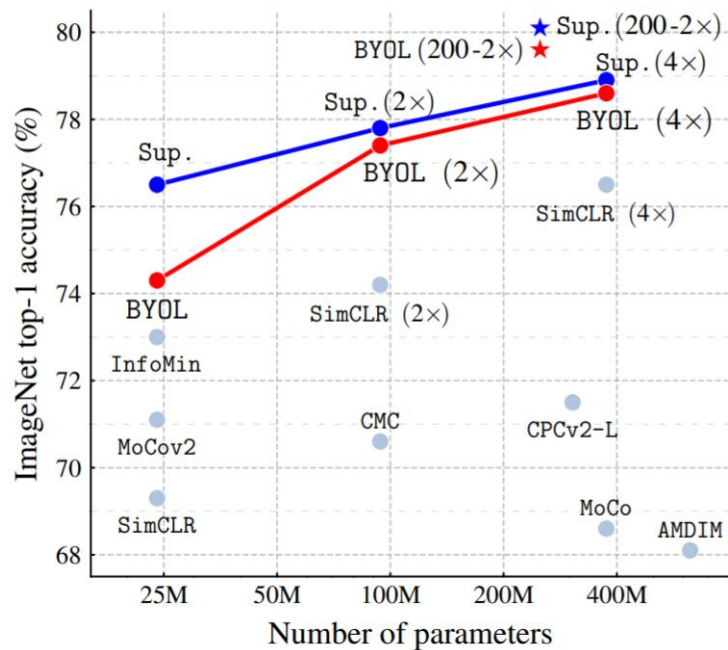
Source: <https://www.freeimages.com/photo/goat-1375679>



Source: <https://www.freeimages.com/photo/sandpiper-going-1373066>

Why Self-supervised Learning?

- Competes with supervised learning



Source: Grill et al., 2020

- Imitates the learning process in humans
- A form of common sense
- Data itself is a signal for learning
- Annotated data is not always available

What is Contrastive Self-supervised Learning?

- A paradigm for self-supervised learning of data representations from **similar and dissimilar pairs**



Source: <https://www.freeimages.com/photo/horse-1393073>



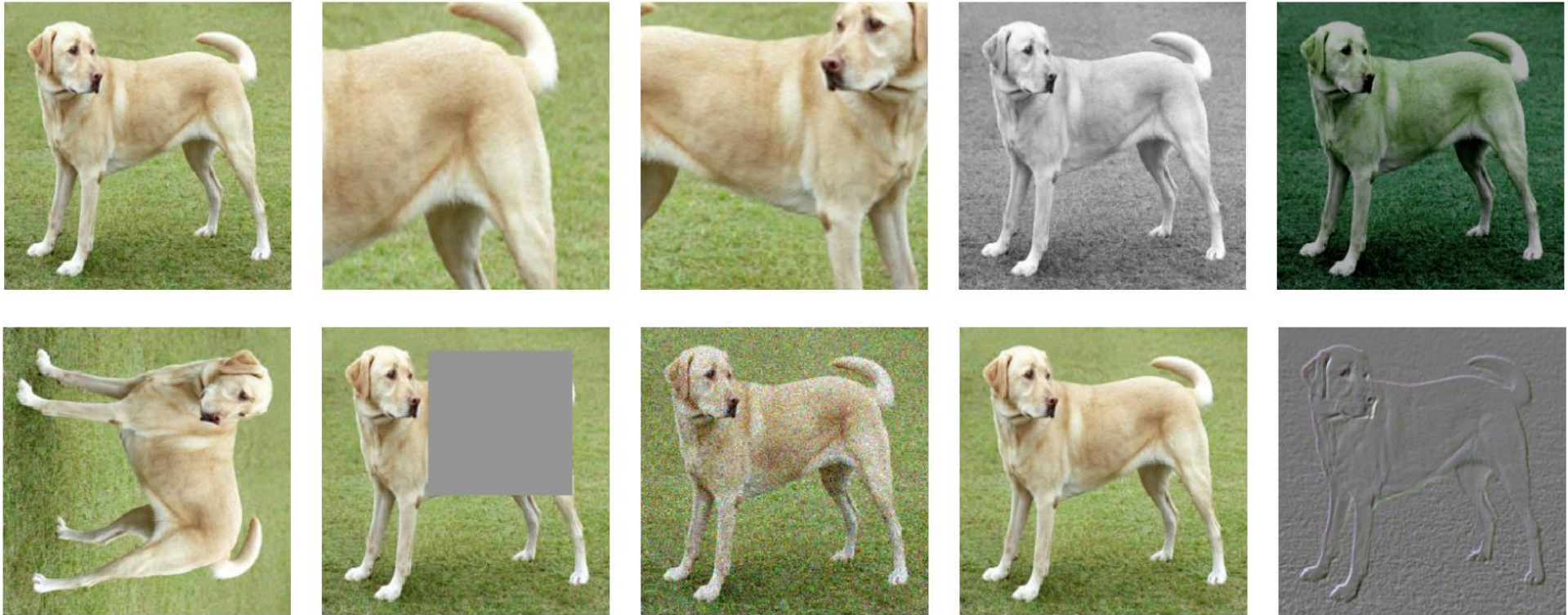
Source: <https://www.freeimages.com/photo/horse-1393073>



Source: <https://www.freeimages.com/photo/goat-1375679>

How to Create Similar Pairs?

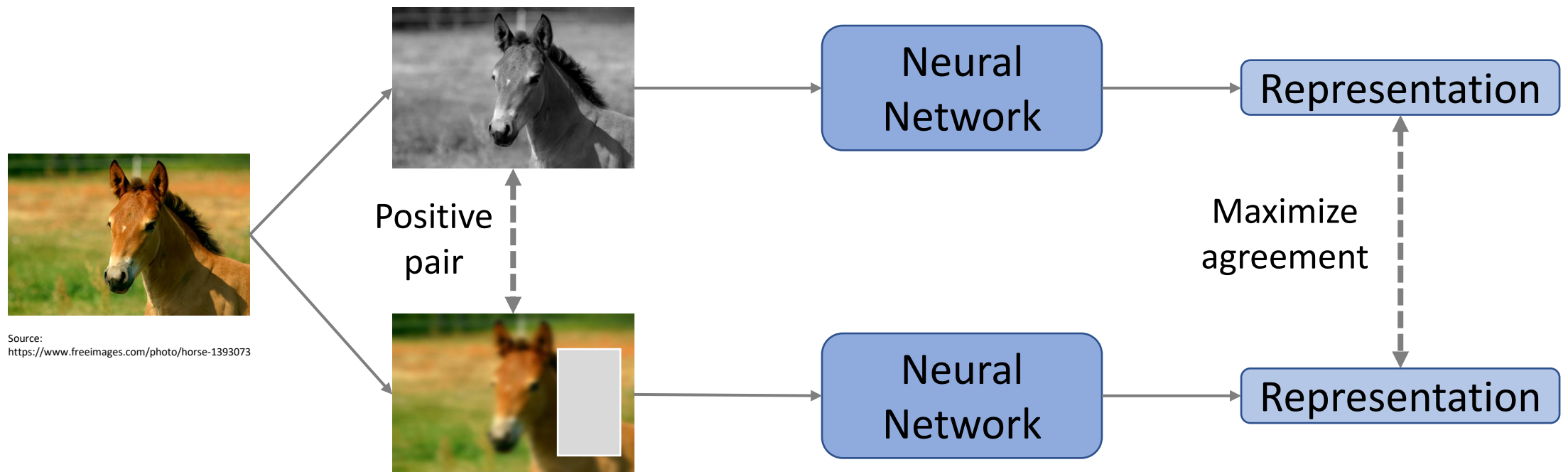
- Data augmentation



Source: <https://arxiv.org/pdf/2002.05709.pdf>

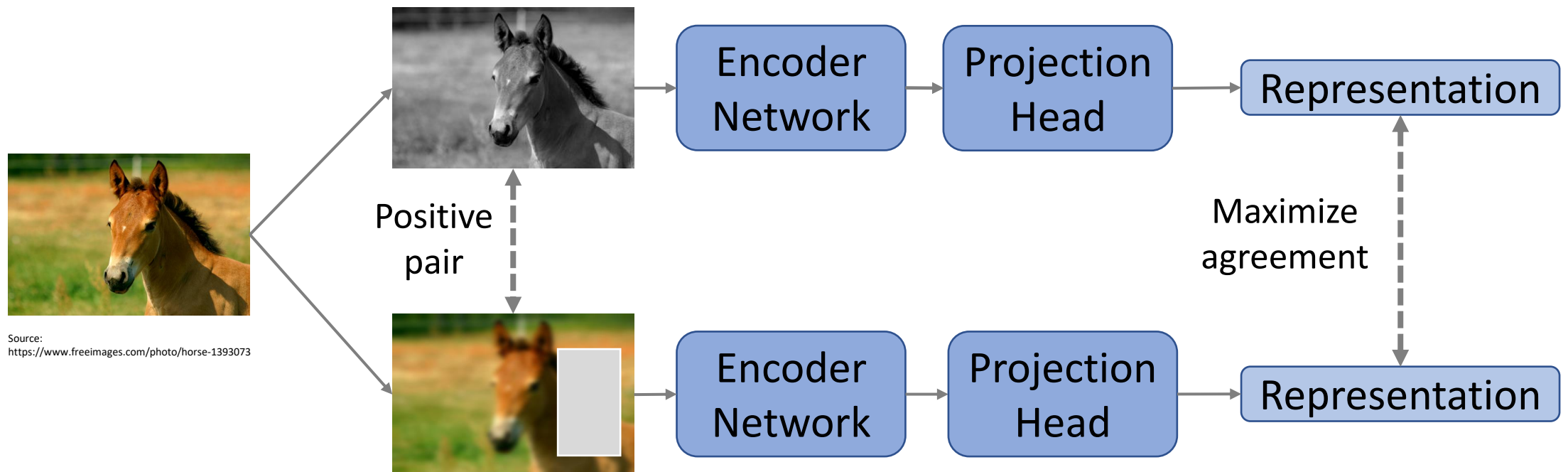
How to: Contrastive Self-supervised Learning?

- Reduce the distance between the representations of positive samples while increasing the distance between negative pairs



How to: Contrastive Self-supervised Learning?

- Reduce the distance between the representations of positive samples while increasing the distance between negative pairs



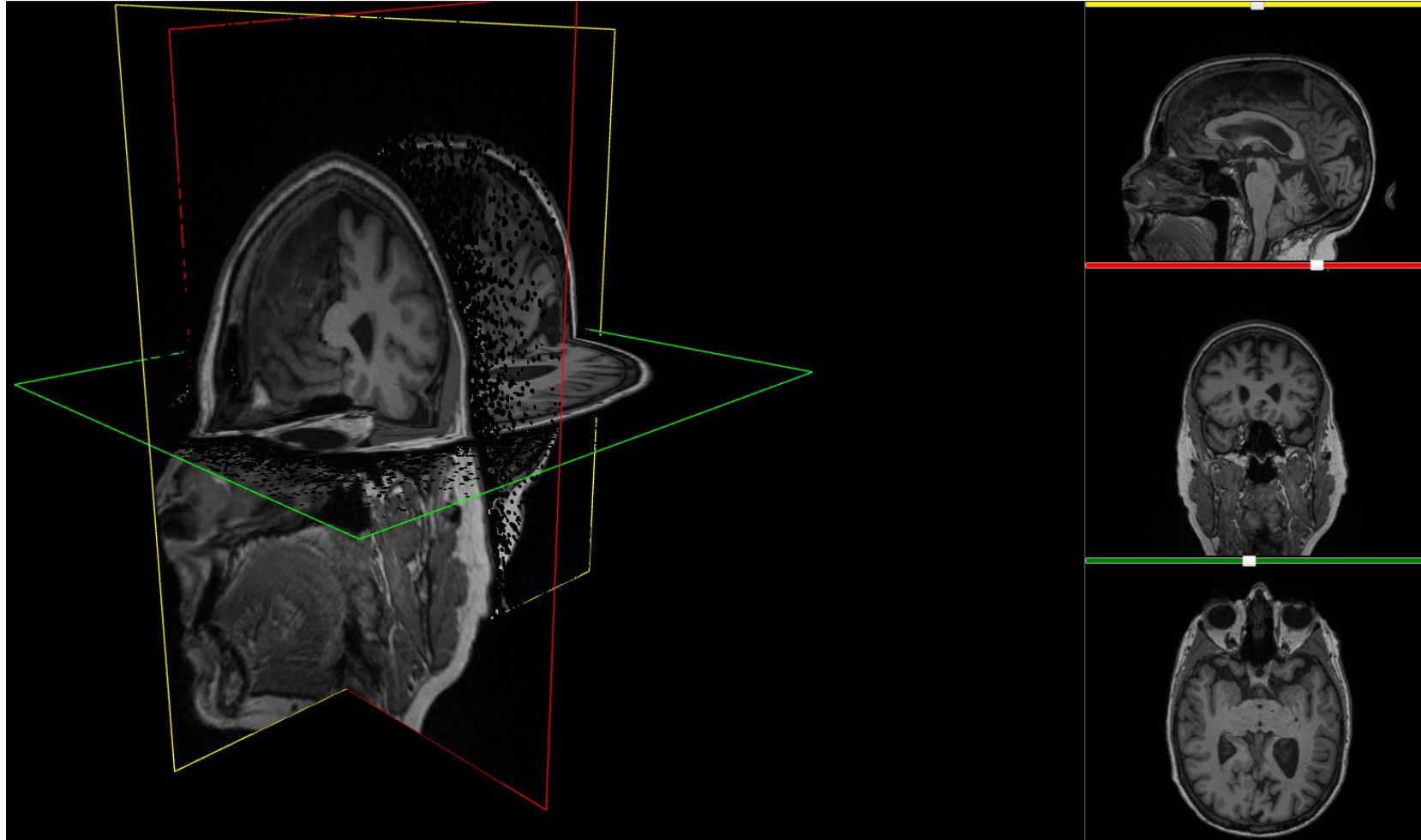
Based on the method by Chen et al., 2020

Outline

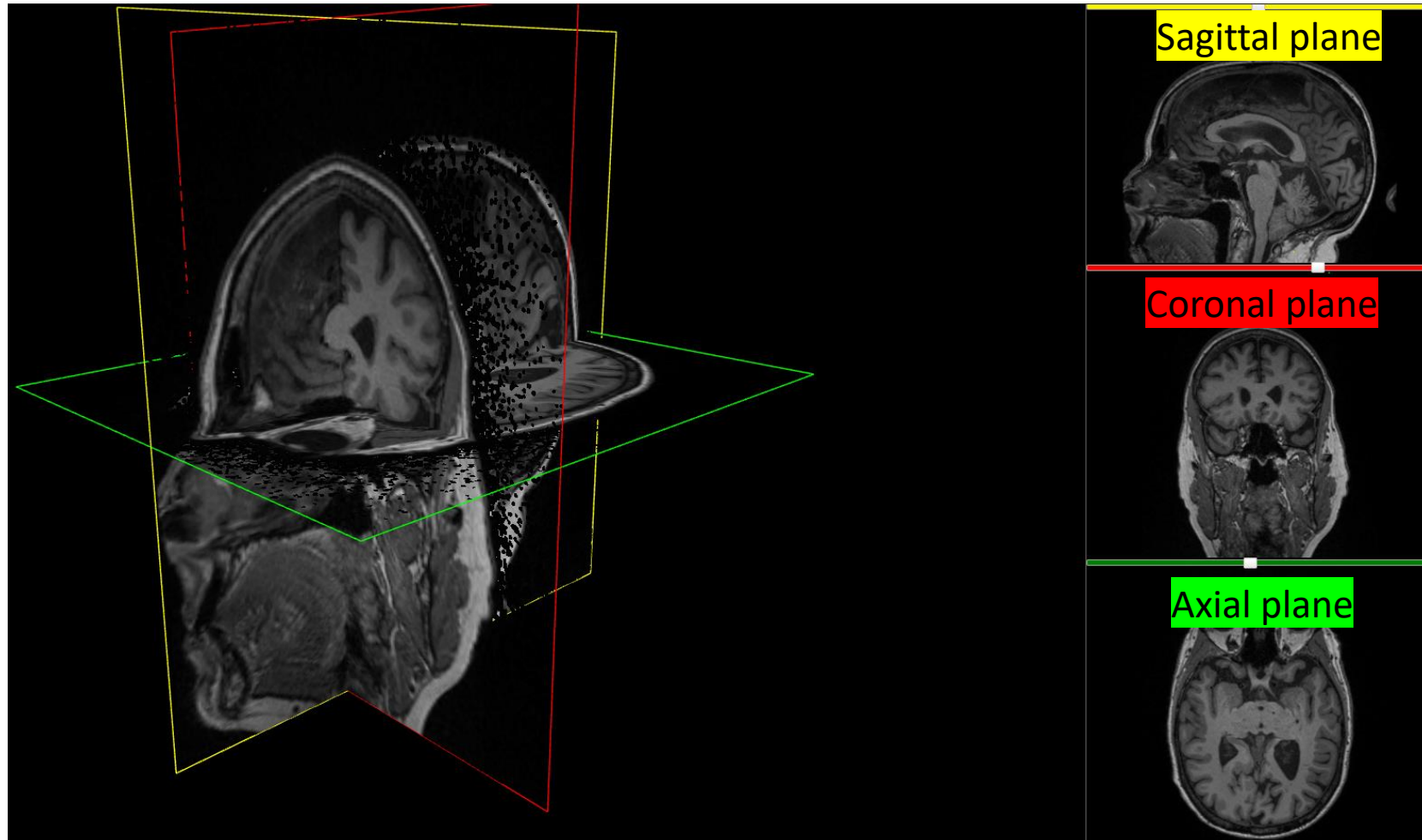
- **Background**
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- Goal
- Architecture
 - Approach
 - Evaluation
- Intermediate Results
- Conclusion & Future Steps



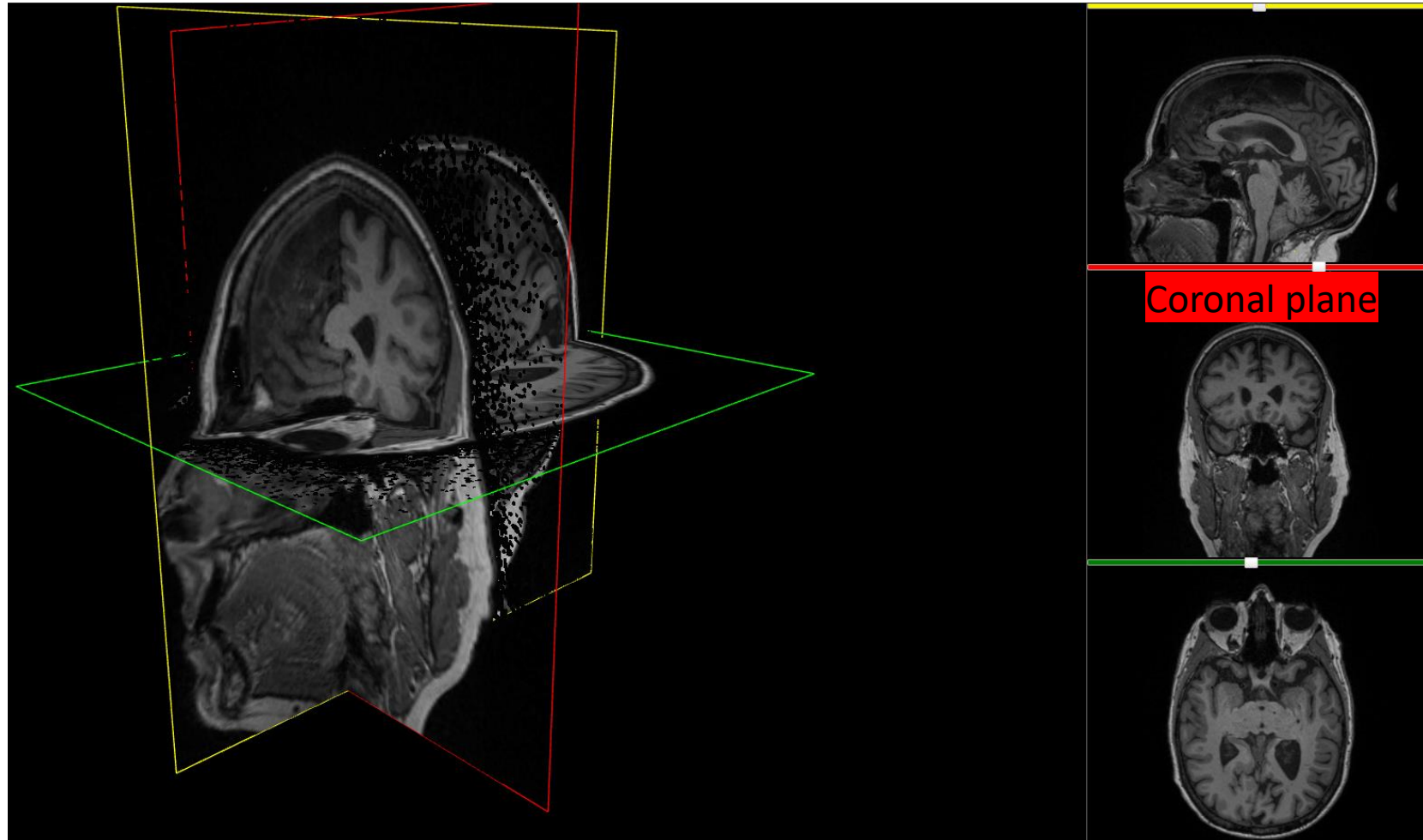
Representation of Data: Structural Magnetic Resonance Imaging



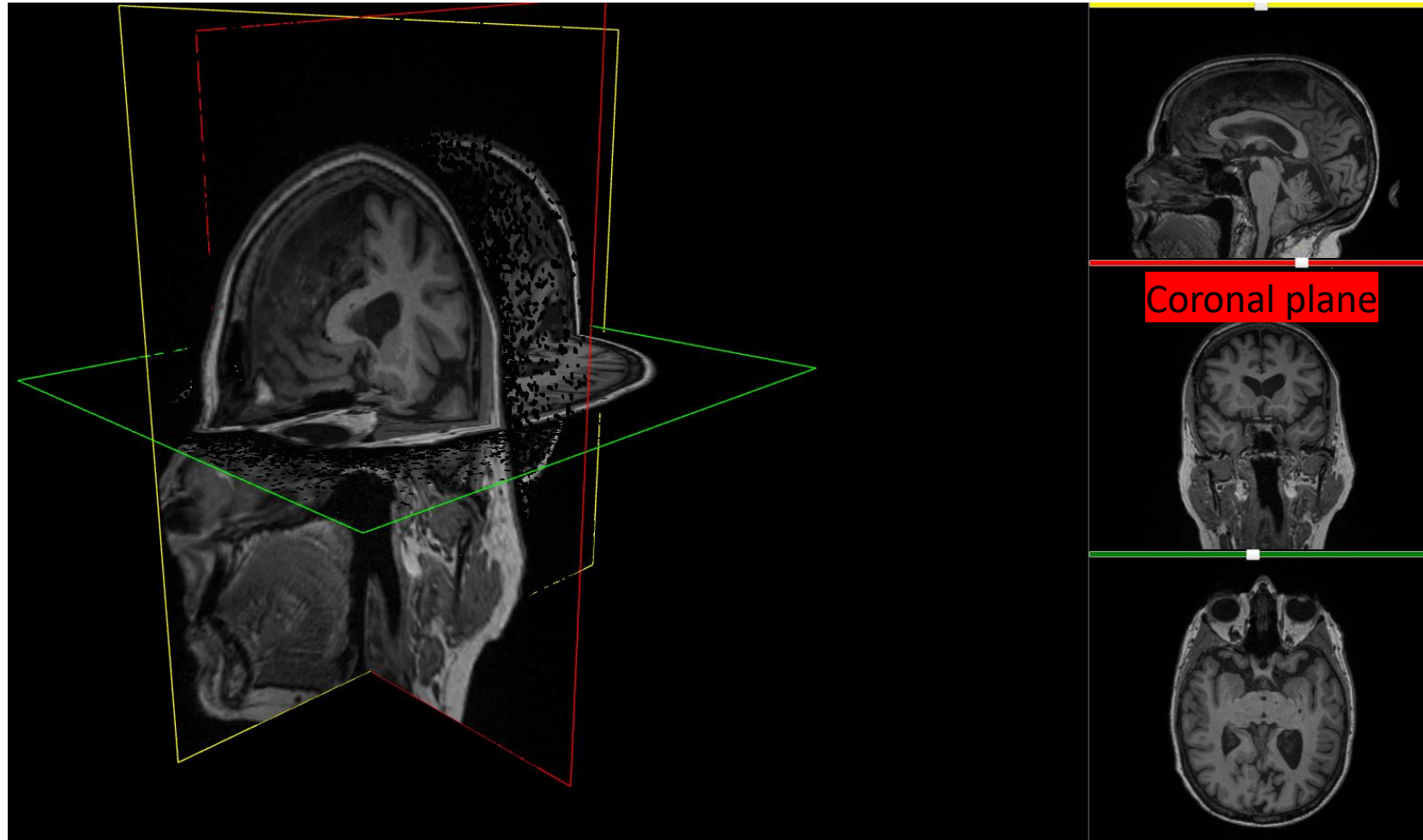
Representation of Data: Structural Magnetic Resonance Imaging



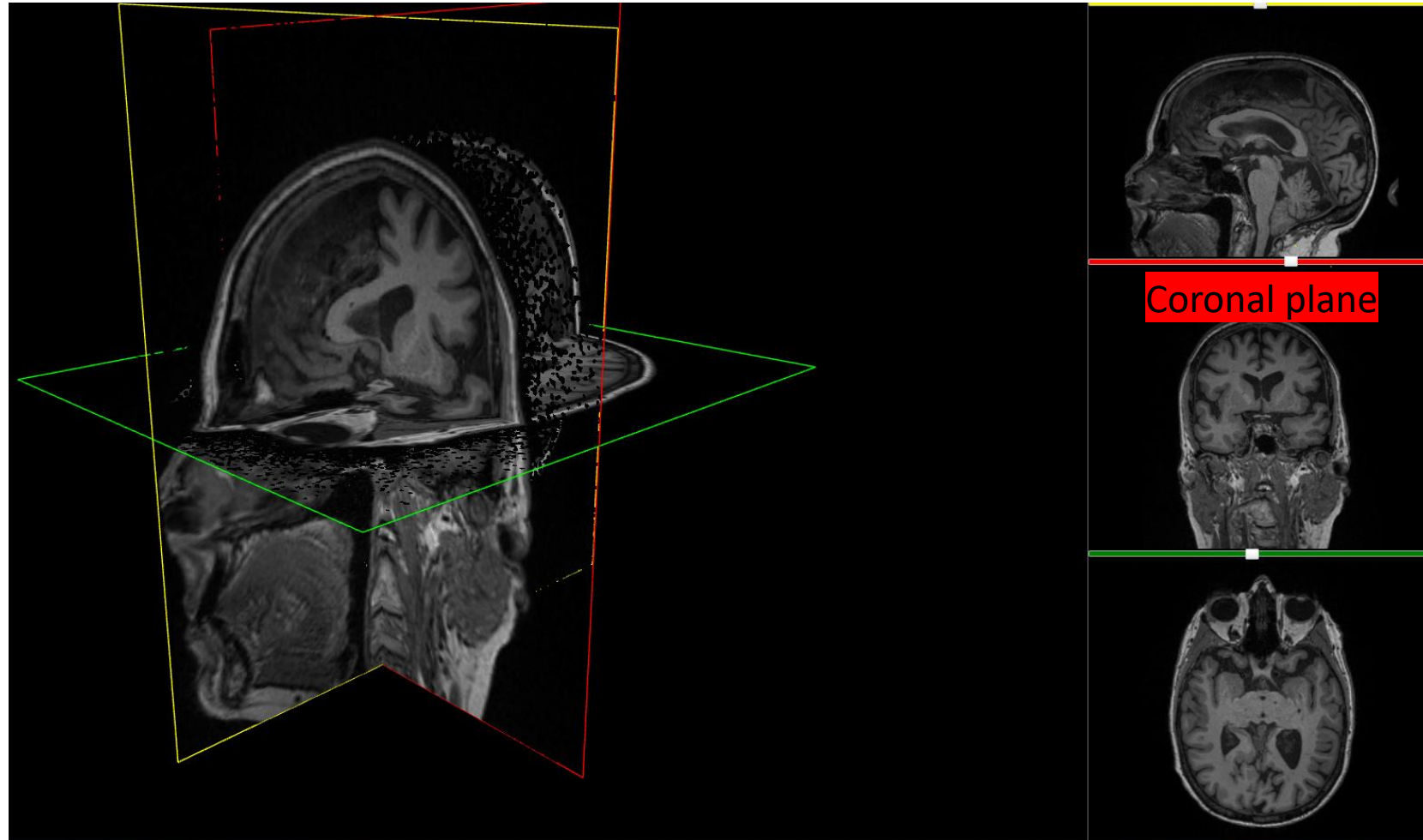
Representation of Data: Structural Magnetic Resonance Imaging



Representation of Data: Structural Magnetic Resonance Imaging



Representation of Data: Structural Magnetic Resonance Imaging

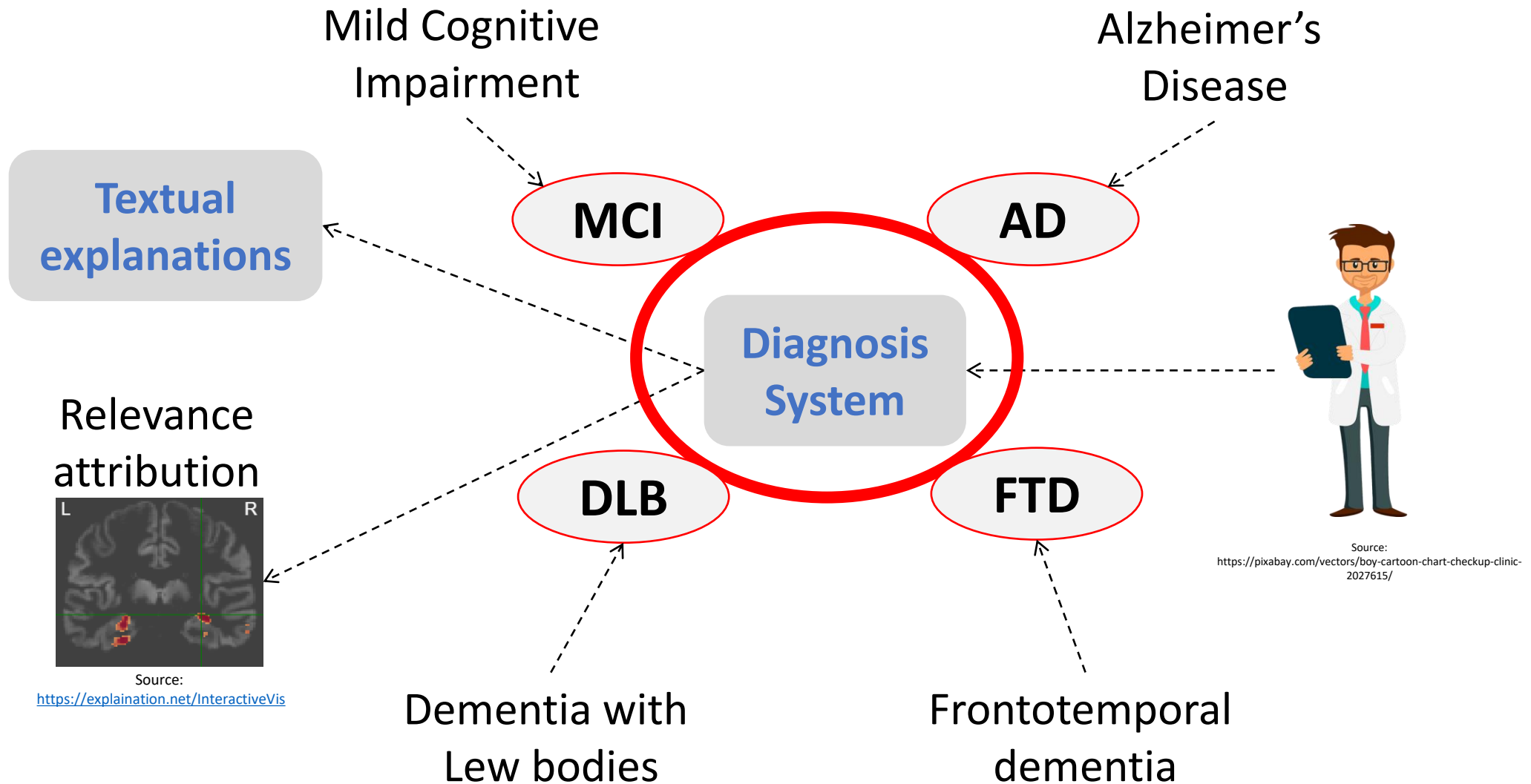


Outline

- Background
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- **Goal**
- Architecture
 - Approach
 - Evaluation
- Intermediate Results
- Conclusion & Future Steps



Goal



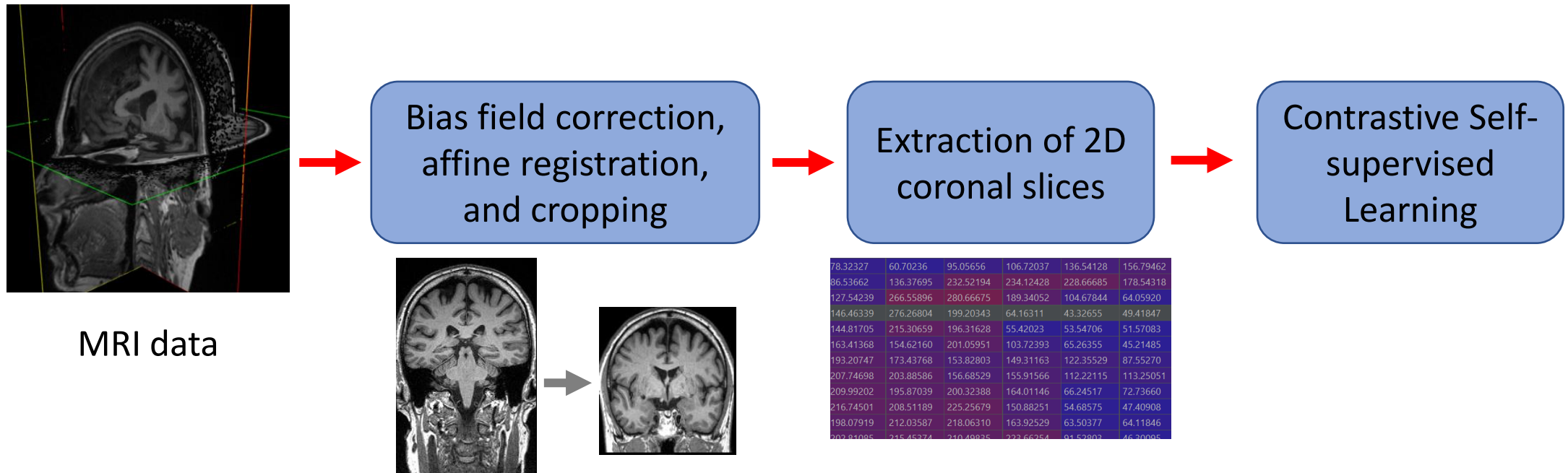
Outline

- Background
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- Goal
- **Architecture**
 - Approach
 - Evaluation
- Intermediate Results
- Conclusion & Future Steps



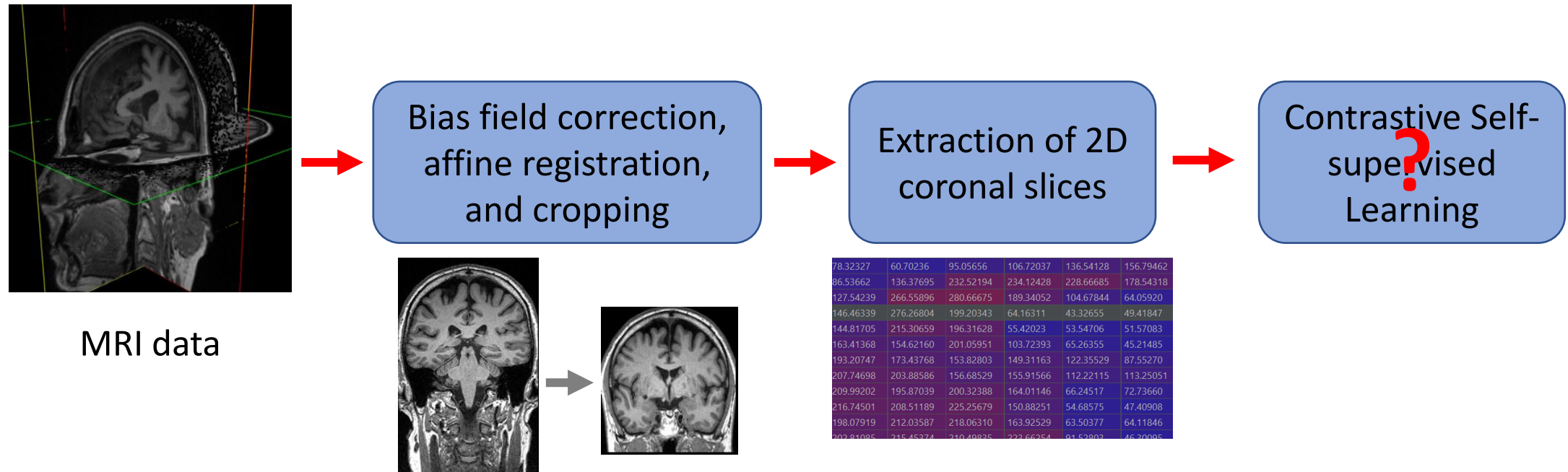
Architecture: Approach

- **Extract** 2D coronal **slices** from MRI data
- Use the slices to **train a neural network** via contrastive learning



Architecture: Approach

- Extract 2D coronal slices from MRI data
- Use the slices to train a neural network via contrastive learning



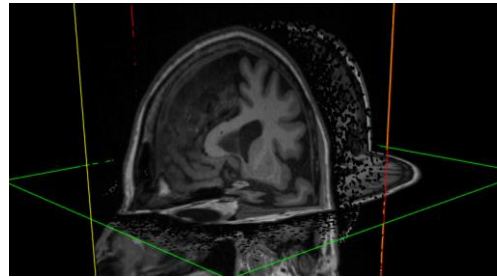
Architecture: Contrastive Self-supervised Learning from MRI Data

- An MRI scan contains **many coronal slices**

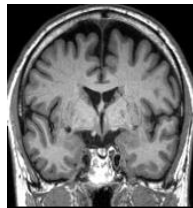


Source:
<https://www.freeimages.com/photo/horse-1393073>

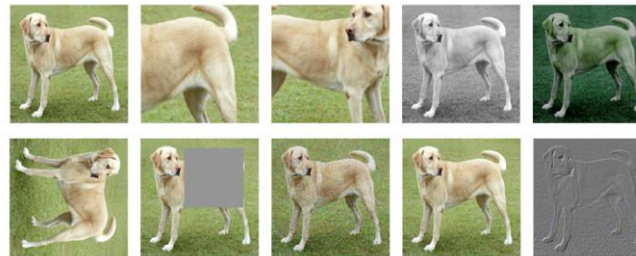
VS



- What **augmentation techniques** can we apply?

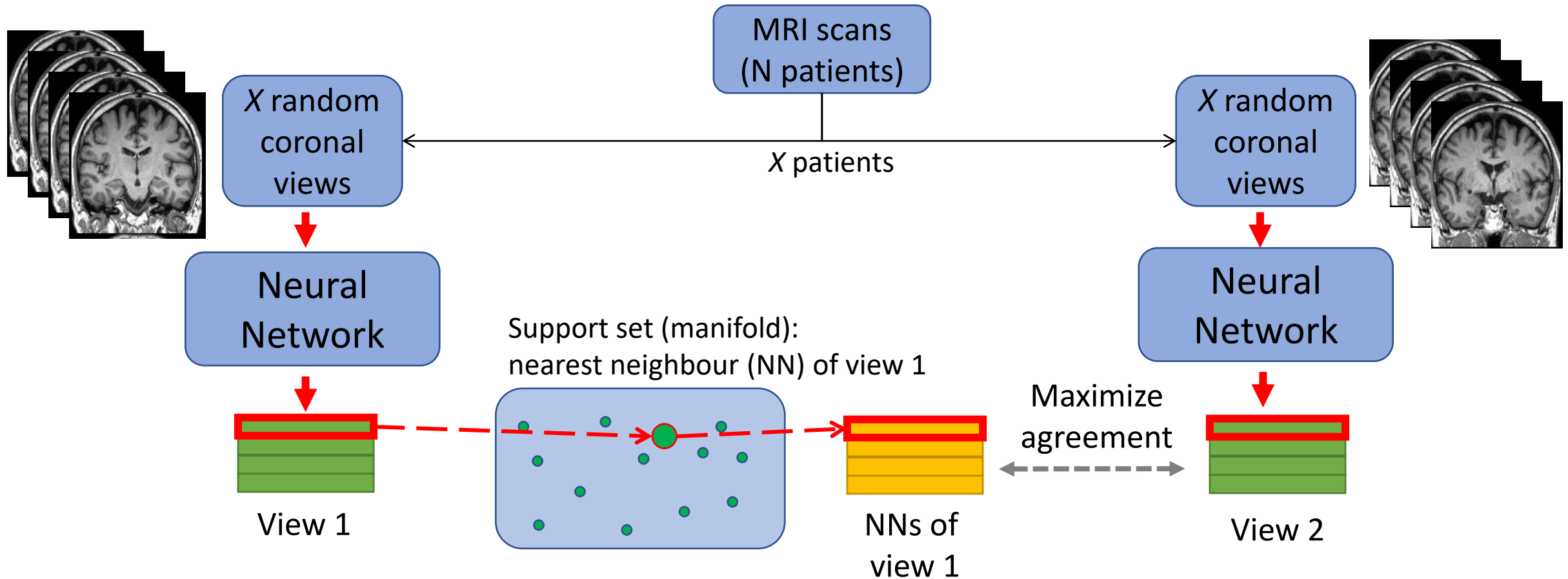


VS



- How do we train a neural network then?

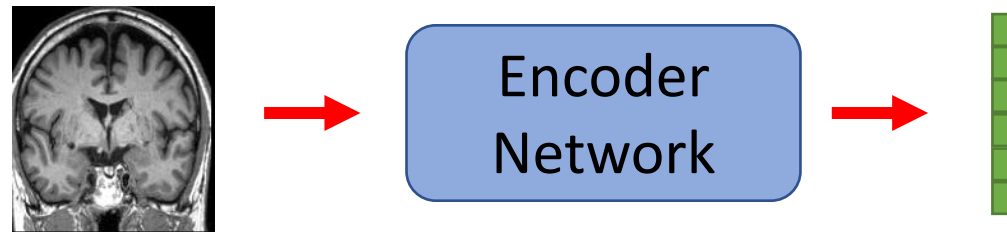
Architecture: Contrastive Self-supervised Learning from MRI Data



Based on the method by Dwibedi et al., 2021

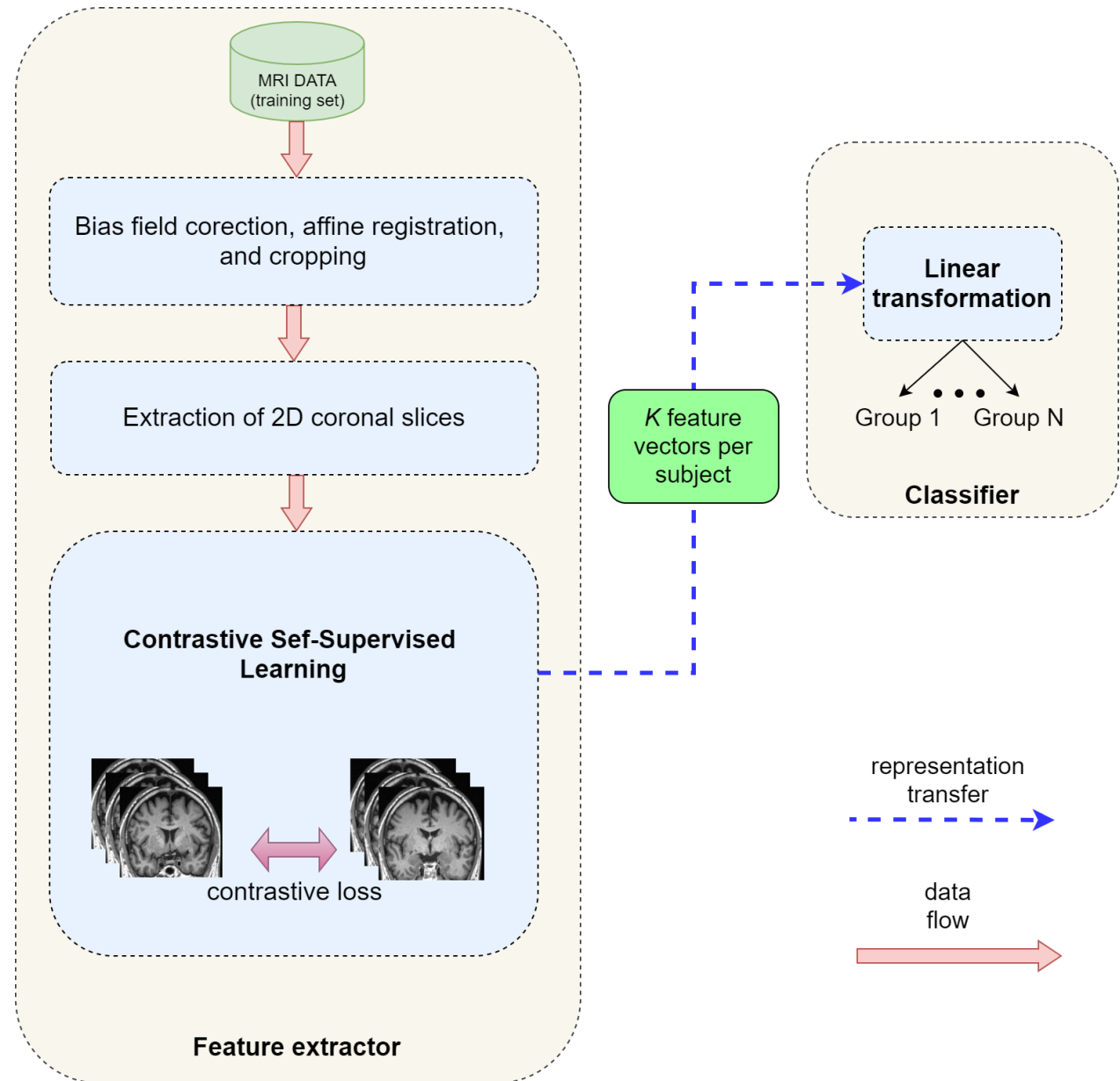
Architecture: Disease Classifier

- An encoder network serves as a **feature extractor** for MRI data (coronal slices)



- **K coronal slices** are queried and respectively **K feature vectors** are extracted for each subject
- These feature vectors are used then to train a **disease classifier** (e.g. control vs patient)

Architecture: Disease Classifier



Outline

- Background
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- Goal
- **Architecture**
 - Approach
 - **Evaluation**
- Intermediate Results
- Conclusion & Future Steps



Planned Evaluation

- MRI Data (**training** and **evaluation**):
 - **ADNI** (Alzheimer's Disease Neuroimaging Initiative), N=2622
 - **NIFD/NIFD** (Neuroimaging in Frontotemporal Dementia), N=346
 - **OASIS** (Open Access Series of Imaging Studies), N=1098
 - **MIRIAD** (Minimal Interval Resonance Imaging in Alzheimer's Disease), N=69
 - **PPMI** (Parkinson's Progression Markers Initiative), N=1841
 - **AIBL** (Australian Imaging, Biomarkers and Lifestyle), N=852
 - **DELCODE** (DZNE Longitudinal Cognitive Impairment and Dementia Study), N=1000
- Quantitative approach
 - Precision & Recall, F_β scores
 - Matthew's Correlation Coefficient
- Qualitative approach



Source:
<https://pixabay.com/vectors/boy-cartoon-chart-checkup-clinic-2027615/>

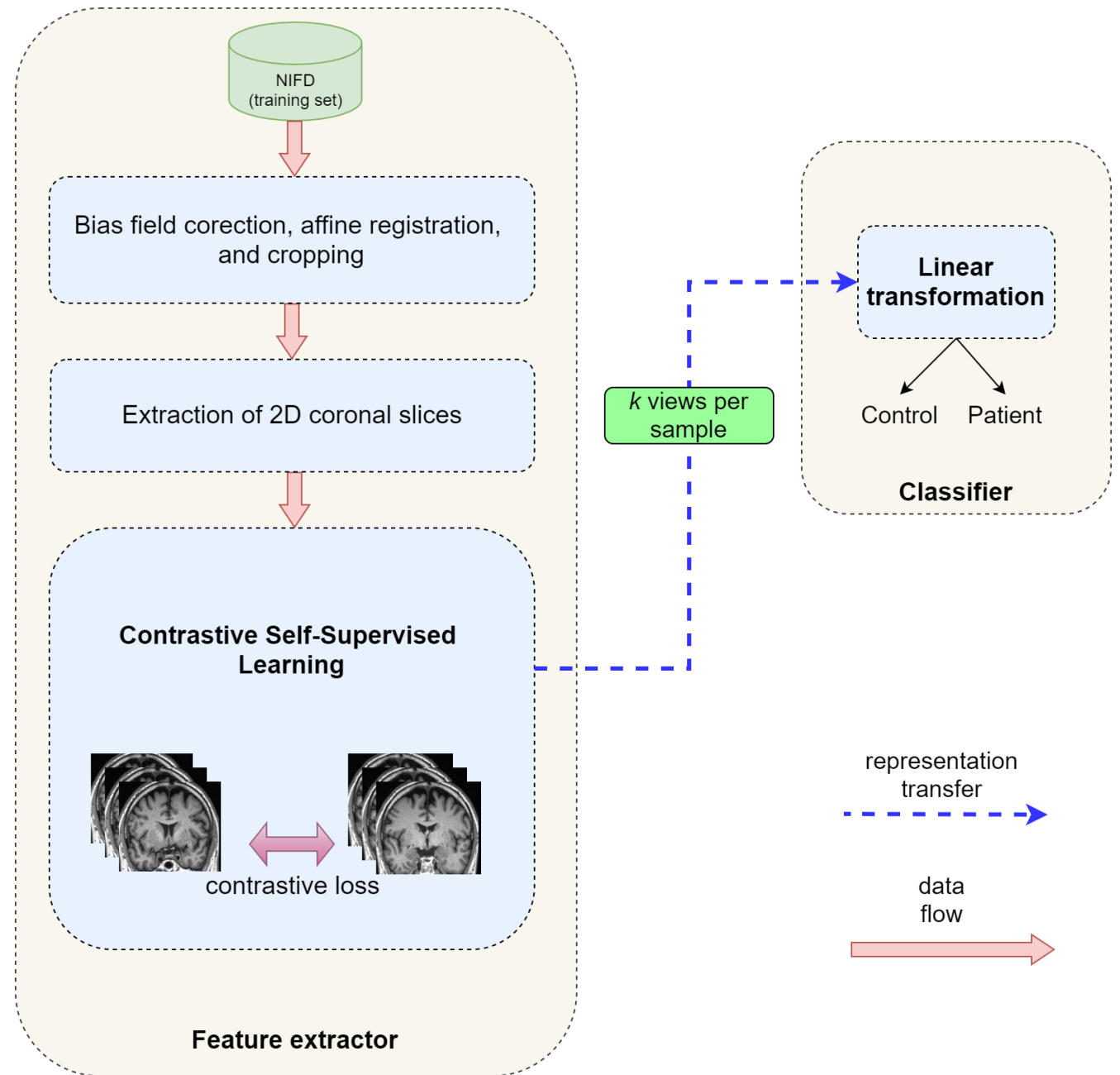
Outline

- Background
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- Goal
- Architecture
 - Approach
 - Evaluation
- **Intermediate Results**
- Conclusion & Future Steps

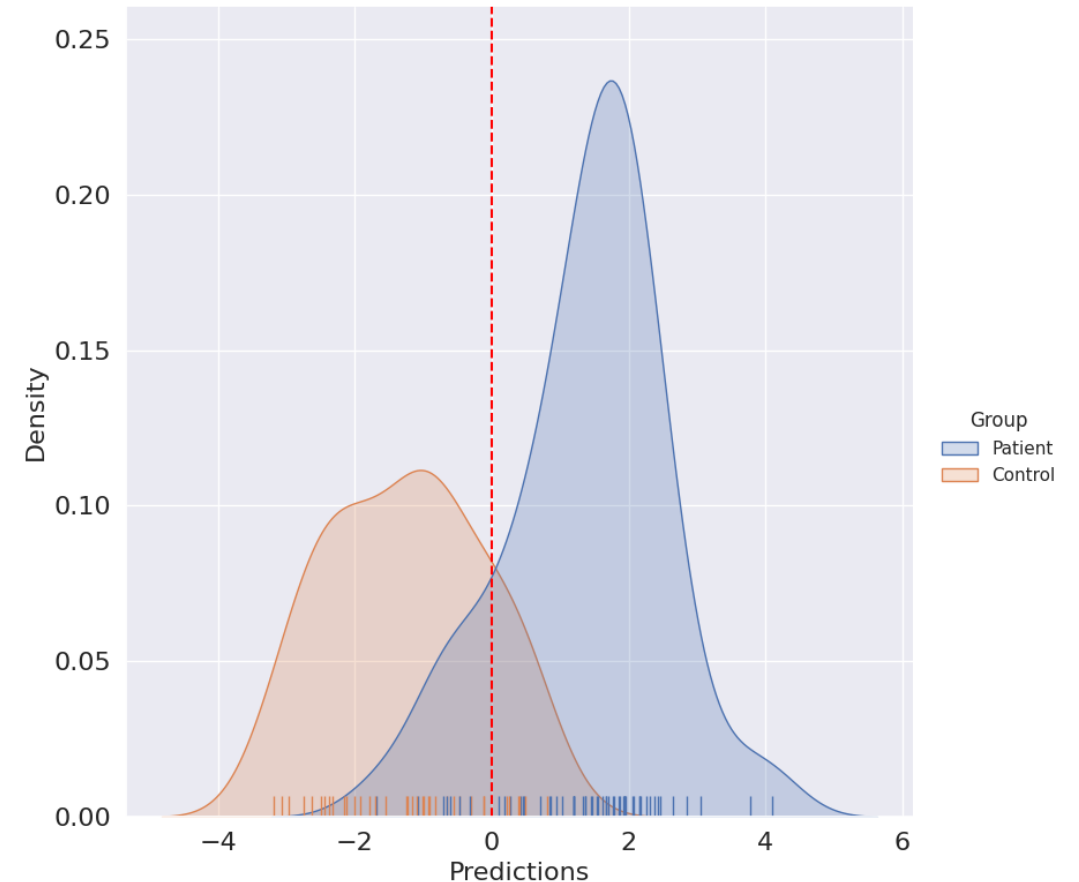
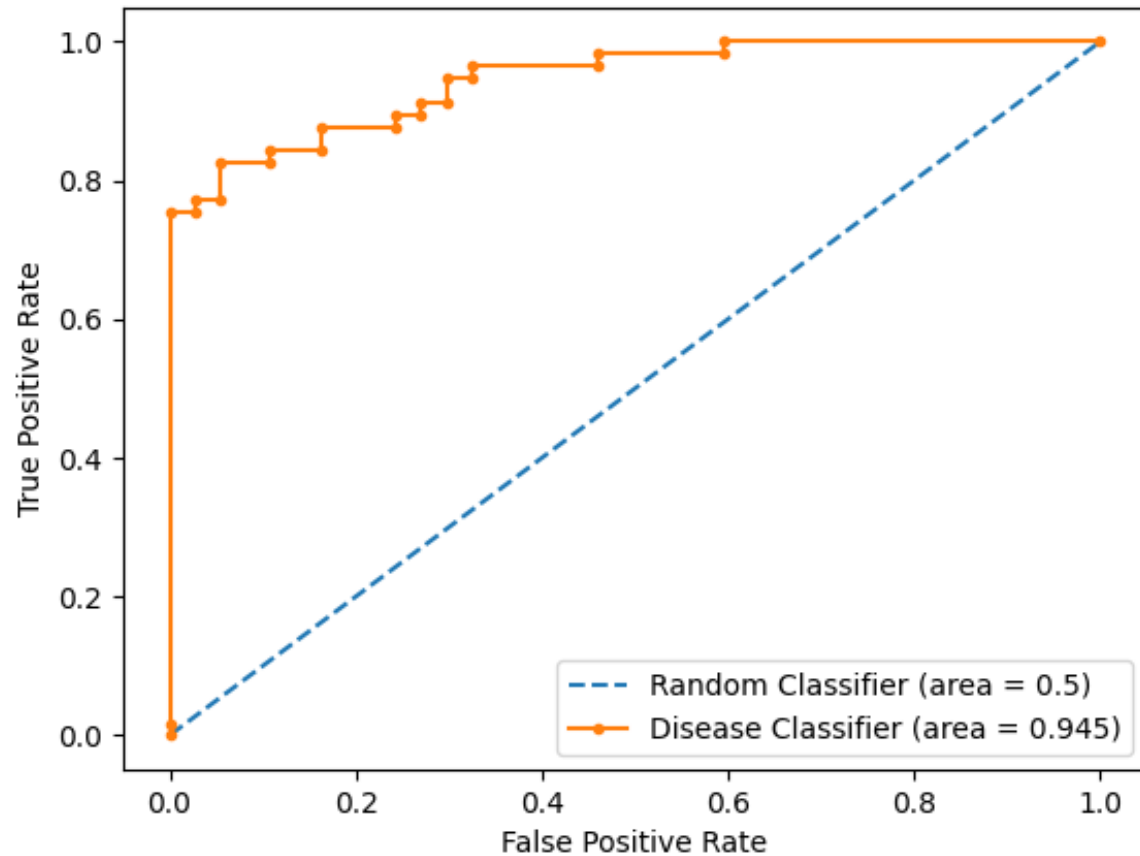


Intermediate Results

- NIFD (Neuroimaging in Frontotemporal Dementia)
 - Training samples: 680
 - Test samples: 94
- Binary classification
 - Control: 379
 - Patient: 395
- Evaluation
 - MCC: ~0,71
 - F1: ~0,88
 - Accuracy: ~0,86



Intermediate Results: Receiver Operating Characteristic and Group Likelihood Estimation



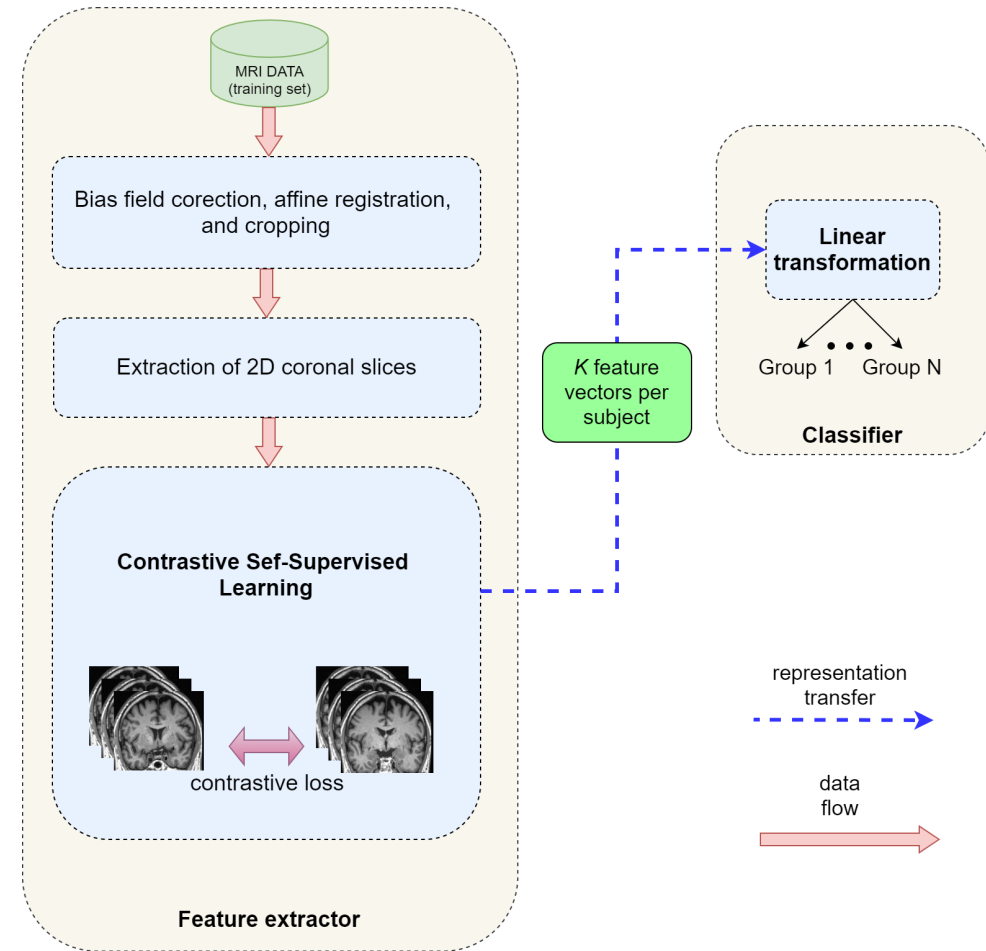
Outline

- Background
 - Contrastive Self-supervised Learning
 - Representation of MRI Data
- Goal
- Architecture
 - Approach
 - Evaluation
- Intermediate Results
- Conclusion & Future Steps



Conclusion & Future Steps

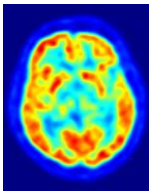
- A system for learning **representations** from MRI data
- **Contrastive learning**
- Aim:
 - A **self-explanatory assistive system** as a support for radiological examination and evaluation
 - Explanations and visualizations as a step to explainable AI (**XAI**)



Conclusion & Future Steps

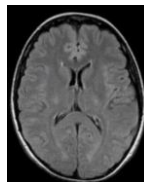
- Multi-Classifier
- Relevance attribution
- Integration of other sources
 - Age, gender, and other tabular data
 - Volumes of the brain regions
- Textual explanations
- Multimodal input

PET



Source:
<https://commons.wikimedia.org/wiki/File:PET-image.jpg>

FLAIR

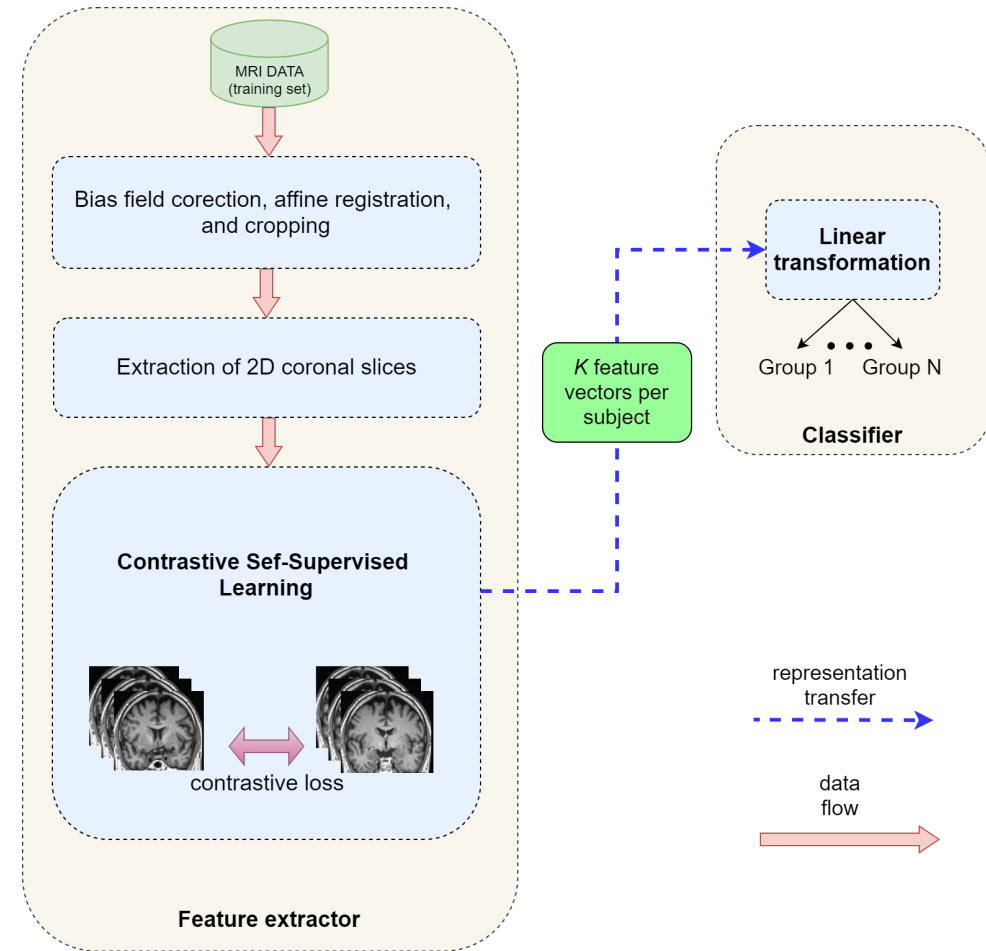


Source:
<https://case.edu/med/neurology/NR/MRI%20Basics.htm>



Source:
<https://pixabay.com/vectors/boy-cartoon-chart-checkup-clinic-2027615/>

- Evaluation by doctors



References

- Dwibedi, D., Aytar, Y., Tompson, J., Sermanet, P., & Zisserman, A. (2021). With a Little Help from My Friends: Nearest-Neighbor Contrastive Learning of Visual Representations. *arXiv e-prints*, *arXiv:2104.14548*
- Grill, J.-B., Strub, F., Altché, F., Tallec, C., Richemond, P. H., Buchatskaya, E., Doersch, C., Pires, B. A., Guo, Z. D., Azar, M. G. & others (2020). Bootstrap your own latent: A new approach to self-supervised learning. *arXiv preprint arXiv:2006.07733*
- Henschel, L., Conjeti, S., Estrada, S., Diers, K., Fischl, B., & Reuter, M. (2020). FastSurfer - A fast and accurate deep learning based neuroimaging pipeline. *NeuroImage*, 219, 117012.
- LaMontagne, P. J., Benzinger, T.L.S., Morris, J.C., Keefe, S., Hornbeck, R., Xiong, C., Grant, E., Hassenstab, J., Moulder, K., Vlassenko, A., Raichle, M.E., Cruchaga, C., & Marcus, D., 2019. *medRxiv*. doi: 10.1101/2019.12.13.19014902
- Malone, I. B., Cash, D., Ridgway, G. R., MacManus, D. G., Ourselin, S., Fox, N. C., & Schott, J. M. (2013). MIRIAD—Public release of a multiple time point Alzheimer's MR imaging dataset. *NeuroImage*, 70, 33–36. doi:10.1016/j.neuroimage.2012.12.044
- Marsland, S., Shapiro, J., & Nehmzow, U. (2002). A self-organising network that grows when required. *Neural Networks*, 15(8-9), 1041-1058. [https://doi.org/10.1016/S0893-6080\(02\)00078-3](https://doi.org/10.1016/S0893-6080(02)00078-3)
- Petersen, R. C., Aisen, P. S., Beckett, L. A., Donohue, M. C., Gamst, A. C., Harvey, D. J., Jack, C. R., Jr, Jagust, W. J., Shaw, L. M., Toga, A. W., Trojanowski, J. Q., & Weiner, M. W. (2010). Alzheimer's Disease Neuroimaging Initiative (ADNI): clinical characterization. *Neurology*, 74(3), 201–209. <https://doi.org/10.1212/WNL.0b013e3181cb3e25>
- Teipel, S., Drzezga, A., Grothe, M. J., Barthel, H., Chételat, G., Schuff, N., Skudlarski, P., Cavedo, E., Frisoni, G. B., Hoffmann, W., Thyrian, J. R., Fox, C., Minoshima, S., Sabri, O., & Fellgiebel, A. (2015). Multimodal imaging in Alzheimer's disease: validity and usefulness for early detection. *The Lancet. Neurology*, 14(10), 1037–1053. [https://doi.org/10.1016/S1474-4422\(15\)00093-9](https://doi.org/10.1016/S1474-4422(15)00093-9)