

Using machine learning in aging neuroscience

By

Christian Johannes Gölz

A thesis presented for the degree of
Doctor rerum naturalium
(Dr. rer. nat)

Paderborn University
Faculty of natural sciences
2022

Contents

Acknowledgement	3
Abstract	3
Figures	4
Tables	5
List of Abbreviations	6
Publications and other scientific contributions	7
1 Introduction	1
2 Theoretical Background	3
2.1 A data driven way to study the aging brain	3
2.2 Machine Learning	3
2.3 Age related reorganization of the brain	4
2.3.1 Contributing Factors	4
2.3.2 Measuring age related changes	5
2.3.3 Neural Datascience	5
2.3.3.1 Dimensionality reduction	5
2.3.3.2 Machine Learning	5
3 Aims and scope	6
4 General methodology	7
5 Publications	8
5.1 Paper 1	8
5.2 Paper 2	8

5.3	Paper 3	8
5.4	Paper 4	8
6	General discussion	9
7	Bibliography	10
8	Statutory Declaration	12

Acknowledgement

Abstract

Aim: Apply data science methods to questions in aging Neuroscience

Methods: Supervised and unsupervised methods in different settings

Results: Novel Data Driven insights

Coclusion: ML rocks!

Figures

Tables

List of Abbreviations

Publications and other scientific contributions

Chapter 1

Introduction

- ML as the next frontier in science
- Open questions in aging neuroscience
- What can ML tell us?
- Age related changes occur at different scales and are manifestet at several levels.
- There is a wide variety in how this changes occur
- Changes are e.g. neural dedifferentiation and compensatory mechanisms (see Reuter Lorenz et al. 2010) and are noticable brain network level and dynamics
- NOTE: Check what EEG studies said about this...
- The idea is to model these changes with tools from datascience to answer questions in aging neuroscience
- First study is about detecting dedifferentiated and compensatory mechanisms with EEG
- Tools used are DMD and Machine learning
- Main idea: Study classification performance as proxy for age related changes in different motor control tasks
- Expertise as possible way of builing a reserve:
- Higher individuality

- Dynamics of dedifferentiation and how do they relate to fitness
- Basic for targeted interventions
- How much and what (relate to Julia)
- Background of ML
- ML as tool
- novel insights
- Problem: Data is multidimensional and we have often limited data
- Solution: Use DMD to reduce Complexity and "model" evolution of signal
- Dynamic Mode Decomposition
- DMD extracts coupled spatio-temporal modes and is able to kind of model the evolution of the signal
- Background + Papers
- Mathematical Formulation
- What can ML tell us?
- ML applied in aging Neuroscience
- Formulating Aims and goals
- Formulation expected outcomes

Chapter 2

Theoretical Background

2.1 A data driven way to study the aging brain

”If our goal as a field is to use data to solve problems, then we need to move away from exclusive dependence on data models and adopt a more diverse set of tools” [1]

Besides classical statistical approaches tools from data science, i.e., machine learning, provide new opportunities to understand multidimensional data in a naturalistic way. Rather than assuming that data is generated by a given stochastic process an algorithmic view treats data mechanisms as unknown [1]. This view is mainly used in the field of data science, which emerged as an independent scientific discipline at the beginning of the 21st century and has adopted concepts from computer science and statistics, especially from the field of machine learning [4]. In the following chapters this framework is outlined in application on aging neuroscience.

2.2 Machine Learning

Machine learning developed as a subbranch of artificial intelligence to enable computers to learn without being explicitly programmed [5]. As such its main purpose is to automatically extract patterns and trends from data or in other words to learn from data [2]

- What is ML and where is it used - ML in the detection of age related changes and why is it of relevance there

2.3 Age related reorganization of the brain

Age related reorganization processes are detectable at the whole body. This is underpinned by multiple interacting biological systems operating on several spatial and temporal scales contributing to the complexity of the phenomenon [3]. At the behavioral level these processes are noticeable in changes in cognitive, motor and sensory functioning [QUELLE]. Aging is one of the biggest risk factors for neurodegenerative diseases such as dementia, including Alzheimer’s disease, as well as Parkinson’s disease making the brain as one of the target systems to study. Patterns of reorganization of the brain are highly individual as they are subject to genetic and environmental influences [QUELLEN]. At the same time, however, overarching, generalizable patterns can be detected [QUELLE]

On a structural level aging has been associated with a reduction in gray matter with an onset early in life

2.3.1 Contributing Factors

Data

The complex interplay of the aforementioned factors leading to the dynamics of brain network changes over the lifetime perspective is not fully understood. However, this is a prerequisite to differentiate healthy from pathological changes and to develop and verify treatments as well as targeted interventions. This requires uncomplicated, easy-to-use, and cost-effective methods and novel analyses to quantify changes in brain organization. Several methods are available to study the brains’ structure and function including functional and structural Magnetic Resonance Imaging (s/fMRI) as well as Magnetoencephalography (MEG) and Electroencephalography (EEG). In addition to cost-intensive imaging techniques such as fMRI, EEG, which meets these criteria, is particularly suitable. Moreover, EEG measures neuronal activity directly with high temporal resolution, which would allow us to gain new insights into the reorganization of brain networks over the lifespan in health and disease. However, it is unclear how the changes described above are reflected in electrophysiological markers. Furthermore, EEG signals are temporally and spatially highly dimensional and have a low signal-to-noise ratio, which makes the detection and visualization of brain networks and their dynamics difficult and requires advanced signal analysis methods. Advanced methods such as methods from the field of artificial intelligence with machine learning, as a subbranch, are of special interest in this context. Methods from supervised and unsupervised machine

learning are possible candidates. In unsupervised machine learning, the goal is to find structure in the data. This includes methods for dimensionality reduction and clustering. Dimensionality reduction for example allows us to describe the structure of high dimensional data in fewer properties [20]. Two common methods in the analysis of neurophysiological data are the principal component analysis (PCA) and independent component analysis (ICA). These allow the detection of spatial patterns in the data that represent the underlying network characteristics of neurophysiological data [8]. In addition, with dynamic mode decomposition (DMD), Brunton, Johnson, et al. [21] apply for the first time a method to electrophysiological data that allows us to map both the spatial and temporal structure of the network structure of neurophysiological data. In supervised machine learning, models are created that can predict a certain outcome based on input data. This method is used to detect neuronal representations of the environment or certain behaviors as well as group memberships and to identify relevant markers [22]. In the context of lifespan changes, complex brain network behavior based on EEG data could be extracted and visualized using dimensionality reduction. Supervised machine learning methods could be used to detect representations of the environment and behavior and to draw conclusions about the differentiation of brain networks. Automatic detection of group membership could further provide new predictors of nervous system states.

2.3.2 Measuring age related changes

2.3.3 Neural Datascience

2.3.3.1 Dimensionality reduction

2.3.3.2 Machine Learning

Chapter 3

Aims and scope

Chapter 4

General methodology

Chapter 5

Publications

5.1 Paper 1

5.2 Paper 2

5.3 Paper 3

5.4 Paper 4

Chapter 6

General discussion

Chapter 7

Bibliography

Bibliography

- [1] Leo Breiman. Statistical modeling: The two cultures (with comments and a rejoinder by the author). *Statistical Science*, 16(3):199–231, 2001.
- [2] Trevor Hastie, Robert Tibshirani, and Jerome Friedman. *The Elements of Statistical Learning. Data Mining, Inference, and Prediction, Second Edition*. Springer-Verlag, New York, second edition, 2009.
- [3] Kathleen M. Mooney, Amy E. Morgan, and Mark T. Mc Auley. Aging and computational systems biology. *WIREs Systems Biology and Medicine*, 8(2):123–139, 2016.
- [4] Erik Lee Nylen and Pascal Wallisch. *Neural Data Science*. Academic Press, 2017.
- [5] A. L. Samuel. Some studies in machine learning using the game of checkers. *IBM Journal of Research and Development*, 3(3):210–229, 1959.

Chapter 8

Statutory Declaration