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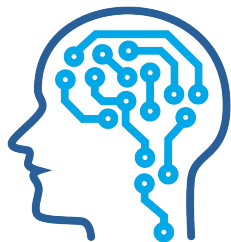
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Leipzig

Medizin ist unsere Berufung.

Medical Data Science

Linking Medicine and Data Science

Leipzig, 02.12.2022



MDS

Medical Data Science

PROGRAM

Medical Data Science - An Overview (~ 20 min)

Prof. Toralf Kirsten

Distributed Analysis using the Personal Health Train (~ 5 min)

Maximilian Jugl

Generation of Synthetic Data (~ 5 min)

Dr. Sina Sadeghi

Privacy-preserving Diagnostic Support for Leukodystrophy Patients (~ 5 min)

Dr. Navid Shekarchizadeh

Predicting the Length of Stay in a Hospital (~ 5 min)

Lars Hempel

Early Detection of Sepsis: The AMPEL Project (~ 5 min)

Dr. Daniel Steinbach

Discussions and Poster Session



Medical Data Science - An Overview

Toralf Kirsten



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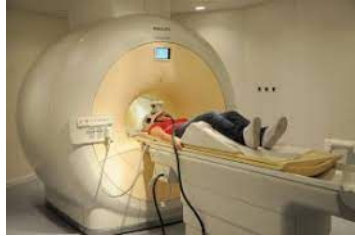
02.12.2022

CHALLENGES IN MEDICINE

- Highly specialized (precision) medicine in an aging society
 - Stratified diagnostics and continued adaptation of care processes
 - Example: Support patients with rare diseases
- Development of new methods in diagnostics and therapies
 - Early diagnostic using available biomarkers
 - Prediction of continued health / care process
- Limited budget

⇒ **Need for efficient use of limited resources**

DATA IS EVERYWHERE



Diagnostics



Intensive Care

Wearables, PROMS, ...



Medical Administration



Clinical Trials, Epi Studies



GOALS

- Capture and manage the right data with the right quality at the right time
- Privacy-preserving data integration and sharing
- Goal-driven data analysis
 - Patient-specific diagnostic support
 - Dedicated to specific medical research questions
 - Administrative reporting

REQUIREMENTS

Infrastructures

Organizational and technical
Infrastructures → FAIR data

Methods

Spectrum allowing to

- capture and integrate data
- prepare and check data
- analyze data

ORGANIZATION AT LEIPZIG UNIVERSITY MEDICINE

**Institute for
Medical Informatics,
Statistics & Epide-
miology**

Medical Departments, Clinics, and Institutions

Junior Research Group
Terminology and
Ontology-based
Phenotyping



MDS
Medical Data Science

Medical Data Science

**Data Integration
Center**

IT Department

Medical Informatics Center (Core Unit)



OUR TEAM



OUR EXPERTISE — OVERVIEW

- **Infrastructure**

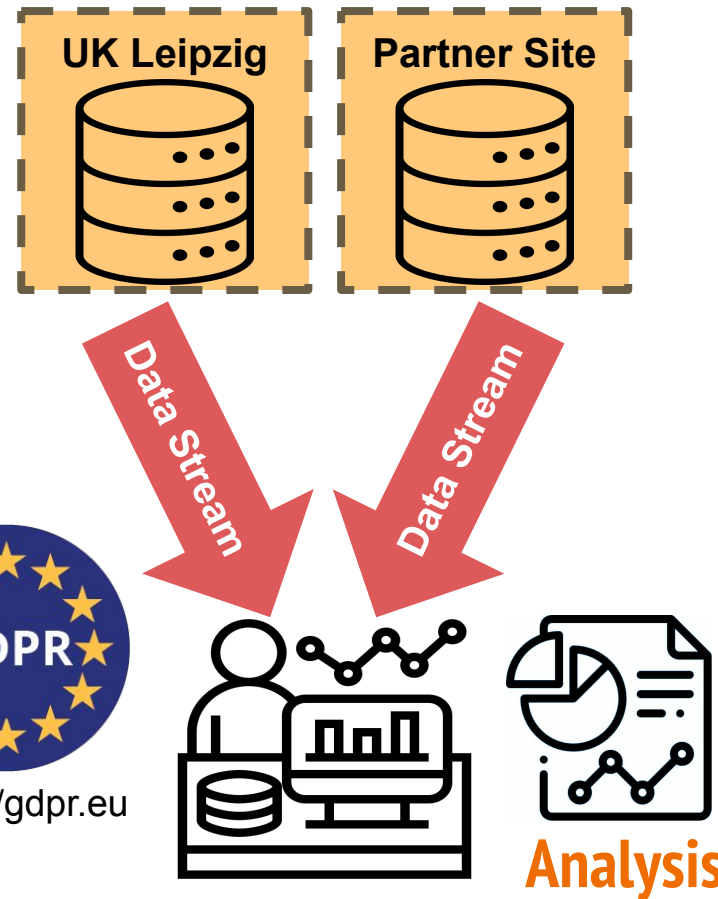
- Enhance data analysis in a distributed mode
- Software applications: patient-specific guiding and diagnostic support

- **Methods**

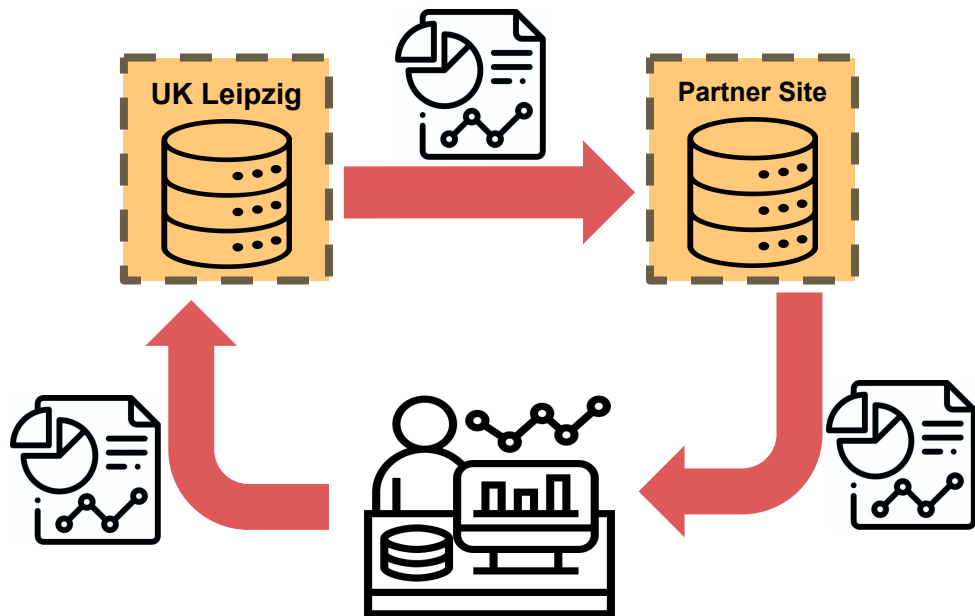
- Artificial-Intelligence-based methods for
 - Early detection, prediction, and diagnosis
 - Identification and segmentation of abnormalities in images
 - Data augmentation, especially for Rare Diseases
- Privacy-preserving methods for data sharing

HOW TO RUN THE ANALYSIS?

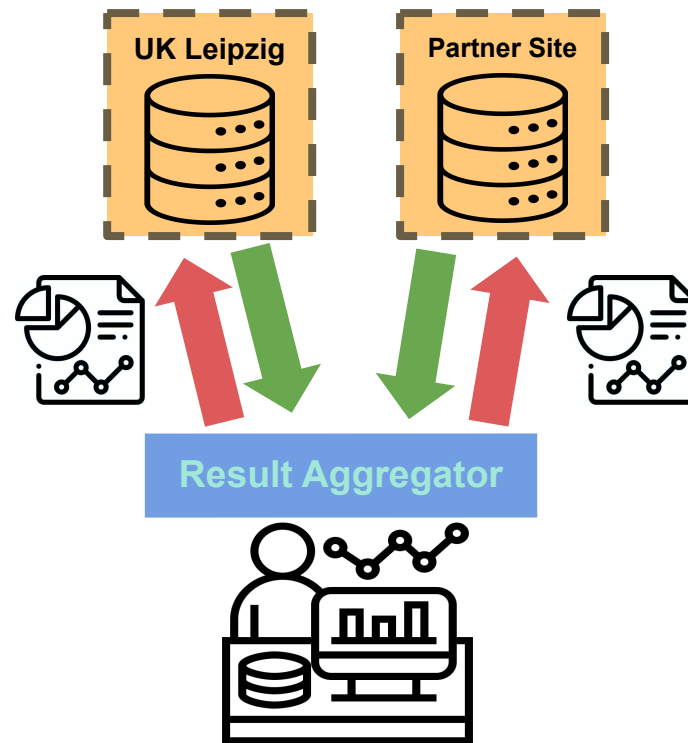
- Centralized (pooled) vs. distributed data analysis
- Influencing factors
 - Number of sites
 - Trust in partner sites
 - Analysis goal and plan
 - Amount and type of data
 - Legal aspect: Consent available?
 - ...



DISTRIBUTED ANALYSIS



Institutional Incremental Learning



Federated/Parallel Learning

PERSONAL HEALTH TRAIN

Publication:

Welten S, Mou Y, Neumann L, Jaberansary M, Ucer YY, Kirsten T, Decker S, Beyan O: A Privacy-Preserving Distributed Analytics Platform for Health Care Data. *Methods of Information in Medicine*. 61(S 01): e1-e11, DOI: 10.1055/s-0041-1740564

- **Key idea:** Bring the analysis to the data
- No direct data access for scientists: Upload the analysis script and run at each involved partner site
- Keep the control over own data at each location
- Strong cooperation with RWTH Aachen, Fraunhofer Institute for Applied Information Technology, Univ. of Cologne, Univ. of Tübingen

⇒ **Pitch Maximilian Jugl**



CHALLENGES IN DISTRIBUTED ANALYTICS

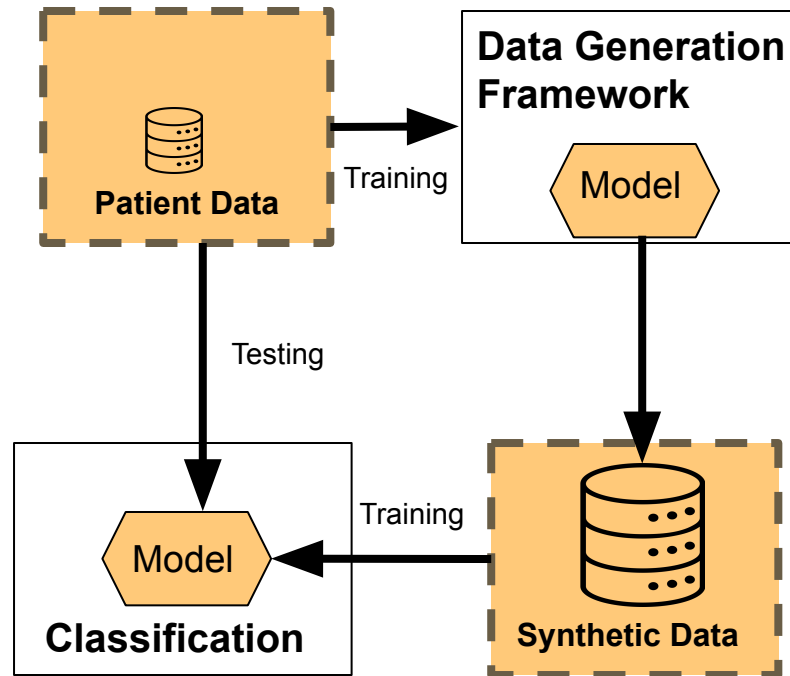


- Infrastructure
 - Computational resources depending on data volume, type and analysis tasks (algorithms) at each location
 - ⇒ **Need to have physical resources or involve further partners**
- Analysis (methods)
 - Impact of routing on final result
 - Catastrophic forgetting: potential drop in accuracy with every new location
 - Unbalanced data: different cohort sizes & class imbalances at different sites, in particular for small data volumes (Rare Diseases)
 - ⇒ **Creation of synthetic data by learning from real world data**

TACKLING SMALL DATA VOLUME BY SYNTHETIC DATA


- **Key Idea:** Use synthetic data to train AI-based models
 - Train a generative model
 - Create synthetic data
 - Train the intended (classification) model using synthetic data
 - Test the intended model by using real world data
- **Our result:** Framework for generating and testing synthetic data

⇒ Pitch Sina Sadeghi




CLASSIFICATION APPROACH FOR POTENTIAL LEUKODYSTROPHY PATIENTS

Close collaboration with local clinicians of the MZEB 



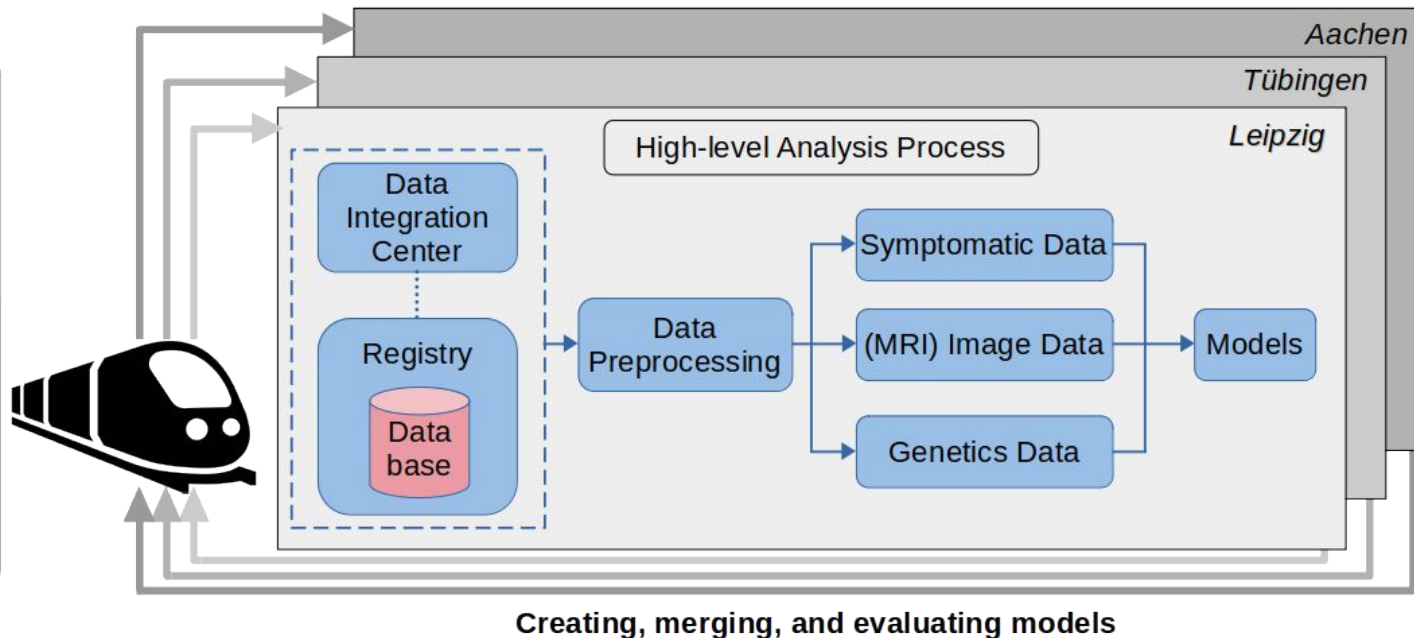
PERSONAL HEALTH TRAIN



PADME

<https://padme-analytics.de>

- ✓ Secure onsite data analysis and result shipping
- ✓ Community-based analysis code review
- ✓ Free of programming languages and data types



⇒ Pitch Navid Shekarchizadeh

Best Poster Award: BMG, Nov. 2022

Designing Diagnostic Support Applications

- **Goal:** Bring research results into practice
- Example: RNFLT-Visualizer (Retinal Nerve Fiber Layer Thickness)
 - Generate new normative data based on a large cohort study (LIFE-Adult)
 - Web-based software application importing and visualizing OCT measurements with normative data



⇒ Poster available



Designing Applications

- Example: Family Manager
 - Capture inheritances of study participants in a pseudonymized way
 - Join other scientific data or consider disease data within a pedigree

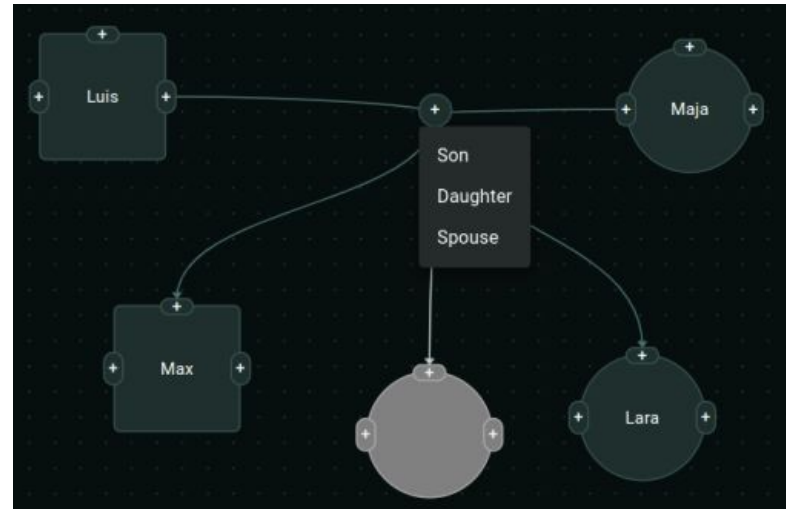


Lara

EDIT

Pseudonym	Lara
Person Type	unknown_patient
Sex	female
Gender	female
Vital Status	alive
Birth Year	2005
Comment	

Family Manager application



SCIENTIFIC PROJECTS WITH FUNDING

- **SMITH (01'2018 - 12'2022, 01'2023 - 12'2026)**
 - Goal: Establishing data integration centers
 - Target: University hospitals
- **NFDI4Health (10'2020 - 09'2025)**
 - Goal: Establishing research infrastructure for study data
 - Target: Clinical trial centers, Large epidemiological studies
- **LEUKO-Expert (10'2020 - 09'2023)**
 - Goal: Guiding support for potential Leukodystrophy patients
 - Target: University Hospitals Leipzig, Tübingen, and Aachen
- **Tag-White (12'2021 - 11'2024)**
 - Goal: Diagnostic support for Leukodystrophy patients
 - Target: Leipzig University Medical Center
- **FAIR - Data Spaces (05'2021 - 04'2024)**
 - Goal: Distributed health data analytics using Gaia-X infrastructure
 - Target: Univ. Leipzig & several partners
- **NUM CODEX+ (01'2022 - 12'2022)**
- **ScaDS.AI (01'2022 - 12'2027)**

Upcoming Projects

- **PATH (12'2022 - 11'2025)**
 - Goal: Secured sharing of health and wellness data
 - Target: Univ. Leipzig & Dresden
- **PrivateAIM (04'2023 - 03'2027)**
 - Goal: Interoperable, secured distributed analytics infrastructure in Germany
 - Target: University Hospitals & other partner sites



OTHER PROJECTS

- **Predicting the length of stay in a hospital** ⇒ Pitch Lars Hempel
 - Goal: Planning support after hospital admission (example: ICU)
 - Target: Leipzig University Medical Center, but could be extended to an overarching project
- **Designing and analysing patient satisfaction (questionnaire-based)**
 - Goal: Analysis of strengths and weakness
 - Target: Leipzig University Medical Center - MZEB
- **DiaClusT**
 - Goal: Find patterns in Diabetes mellitus type 2 patients for a better and dedicated treatment
 - Target: Leipzig University Medical Center (extended to all University Hospitals planned)
- **Privacy-preserving record linkage**
 - Goal: Establishing an infrastructure for record linkage as part of the PHT
 - Target: All University Medical Centers and other partners (e.g., registries)

Distributed Analysis: The Personal Health Train

Maximilian Jugl



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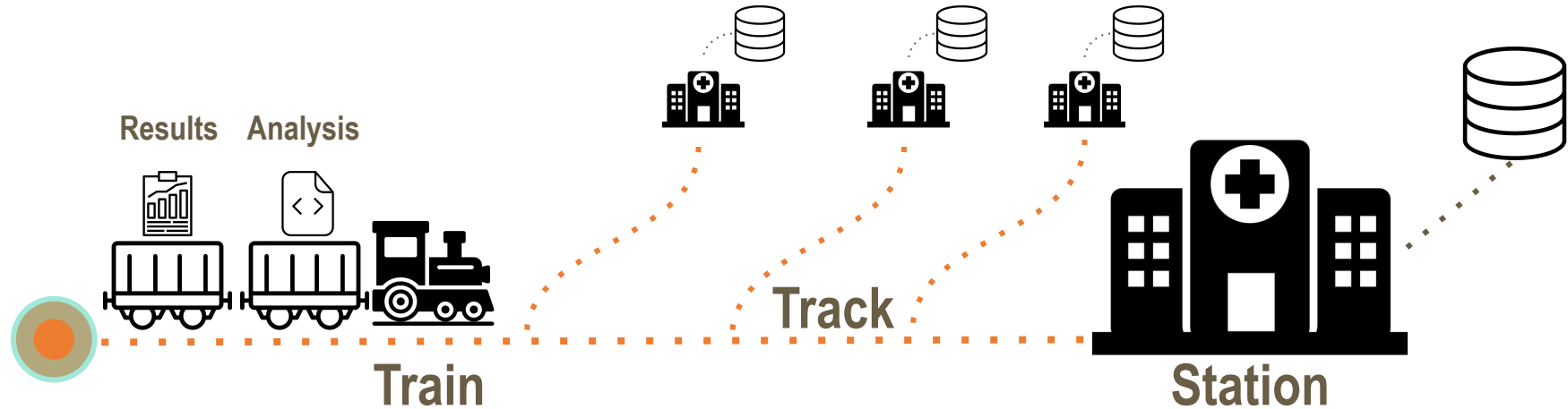
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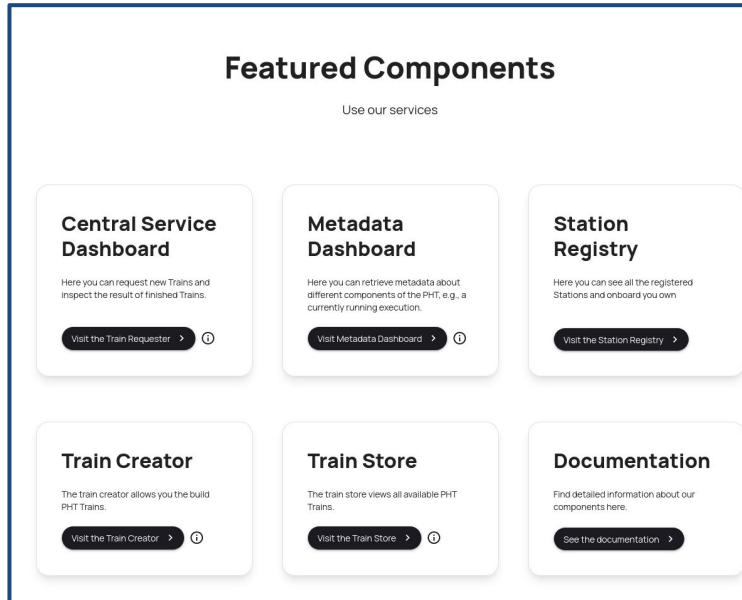
PERSONAL HEALTH TRAIN



- **Paradigm:** Bring the analysis to the data
- The data remains under the control of the data holders (**Station**)
- Analysis travels between stations to update the results (**Train**)



PERSONAL HEALTH TRAIN — PADME



- PHT implementations are in active development (e.g. PADME, VANTAGE6)
- Web portal with access to different components for running distributed analysis tasks



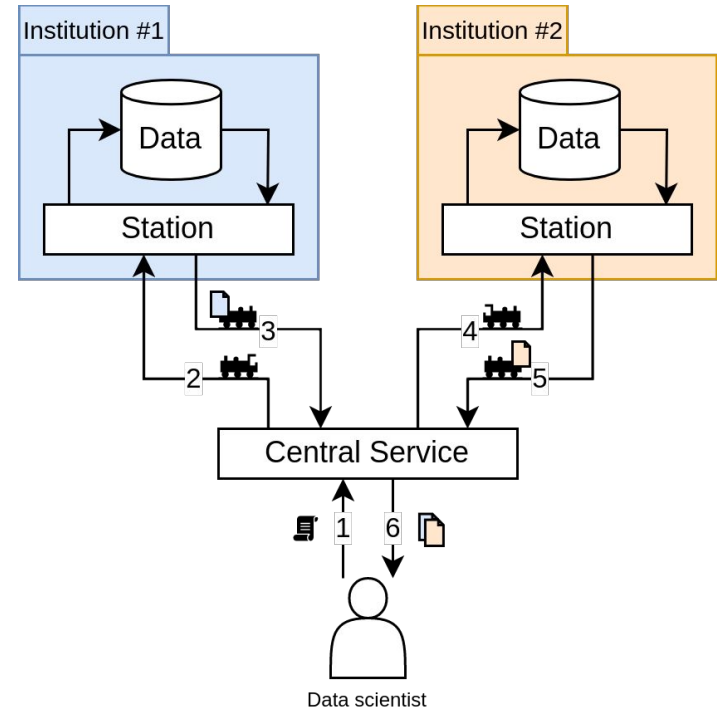
padme-analytics.de

Publication:

Welten, S., Mou, Y., Neumann, L., Jaberansary, M., Ucer, Y. Y., Kirsten, T., ... & Beyan, O. (2022). A **Privacy-Preserving Distributed Analytics Platform for Health Care Data**. *Methods of Information in Medicine*.

PERSONAL HEALTH TRAIN — Procedure

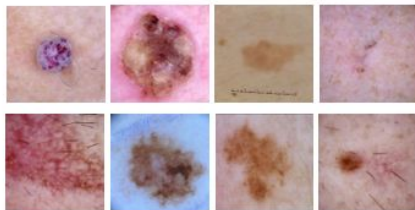
- Data scientist uploads analysis script to **central service** (CS)
- CS constructs **train class**
- Data scientist selects train class and target stations
- CS sends train to selected stations
- Station administrators provide access to relevant resources to train
- Train aggregates results and provides them to data scientist when back at CS



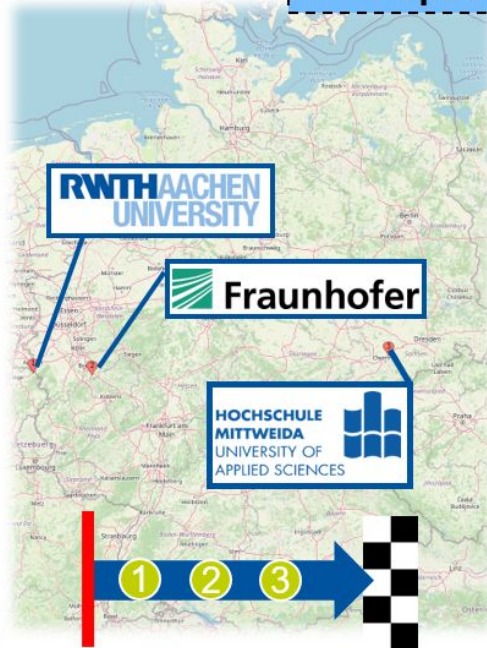
PHT APPLICATION SCENARIO

Dataset & Task

- ISIC 2019¹ Data set
- Classify dermoscopic images
- **Eight different diagnostic categories:**



Participants



Results

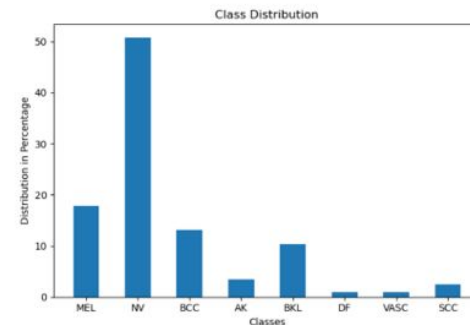
- **General: Comp. to central model**
- **Some metrics:**

Central vs. PHT

75.40 vs. 71.83% (accuracy)

69.22 vs. 63.35% (recall)

- **In-depth statistical analysis:**



Study setup

Collaboration

Outcome

¹ <https://challenge.isic-archive.com/landing/2019/> (Open-Source)

OTHER APPLICATION SCENARIOS

- Lesion detection for Leukodystrophies ⇒ **Poster available**
 - Design of an image synthesis and processing pipeline
 - Train is used for classification of annotated MRIs
- Privacy-preserving record linkage ⇒ **Poster available**
 - Integration of web services for PPRL into the PHT
 - Validation on synthetic data, real-world use case coming soon™
- Model training on skin lesions from ISIC dataset with 3 stations
- Model training on Breast Cancer Wisconsin dataset with 6 stations

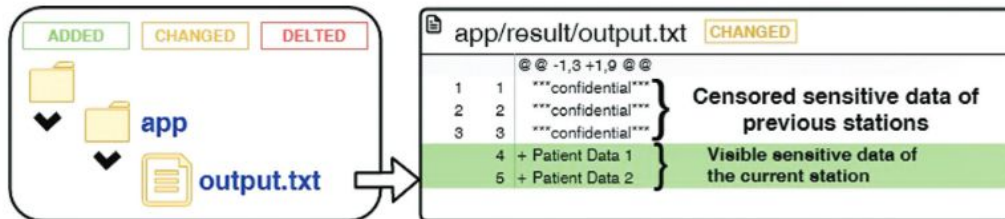
Publications:

- 1) Mou, Y.; Welten, S.; Jaberansary, M.; Ucer Yediel, Y.; Kirsten, T.; Decker, S.; Beyan, O.; **Distributed Skin Lesion Analysis Across Decentralised Data Sources**. Stud Health Technol Inform. 2021.
- 2) Welten, Sascha; Hempel, Lars; Abedi, Masoud; Mou, Yongli; Jaberansary, Mehrshad; Neumann, Laurenz; Weber, Sven; Tahar, Kais; Yediel, Yeliz; Löbe, Matthias; Decker, Stefan; Beyan, Oya; Kirsten, Toralf; **Multi-Institutional Breast Cancer Detection Using a Secure On-Boarding Service for Distributed Analytics**. Applied Sciences 12. 4336. 2022.



SECURITY IN THE PERSONAL HEALTH TRAIN

- Trains are encrypted between stations using public-key cryptography
- Station administrators are in control of the analysis execution
 - Define access to shared resources with train
 - Reject malicious trains
 - Inspect results after analysis execution
- Only analysis results leave the station



Publication:

Welten, S., Mou, Y., Neumann, L., Jaberansary, M., Ucer, Y. Y., Kirsten, T., ... & Beyan, O. (2022). **A Privacy-Preserving Distributed Analytics Platform for Health Care Data**. *Methods of Information in Medicine*.

Generation of Synthetic Data

Sina Sadeghi



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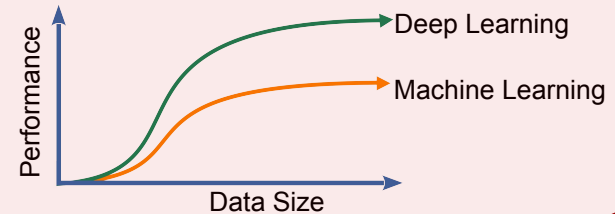
02.12.2022

Medical Data Scarcity

- Patient Privacy
(General Data Protection Regulation)
- Rare Diseases
(with limited available data)

Artificial Intelligence Models

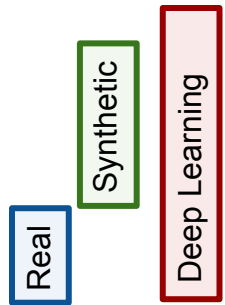
- require a large amount of data



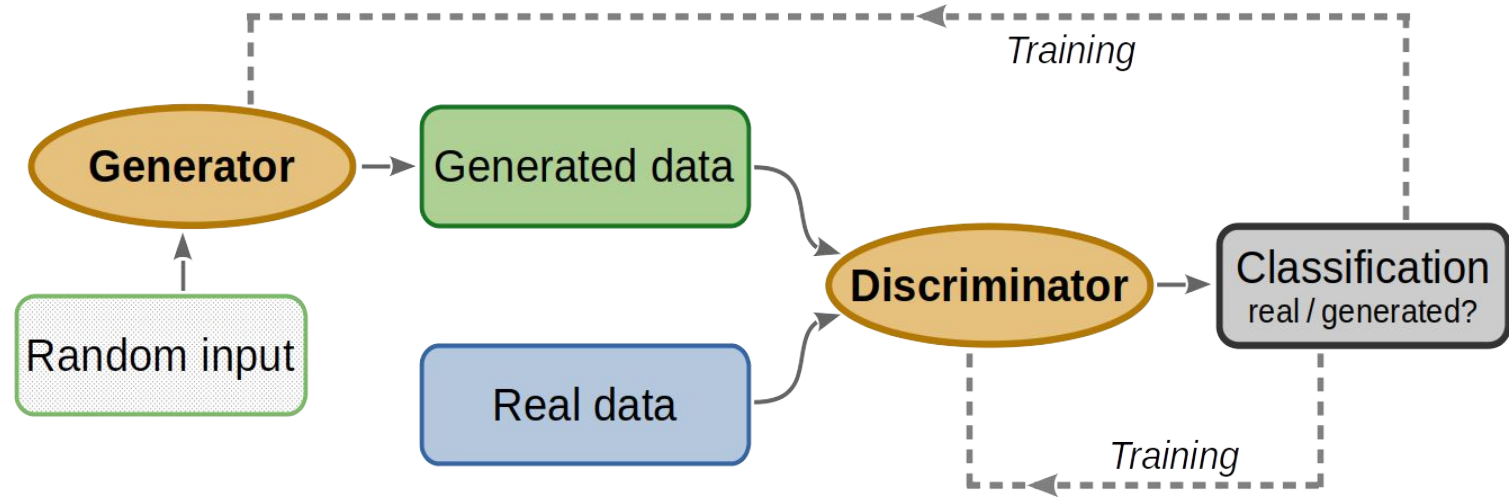
Generating Synthetic Data

- without reference to real patients
- improve data analysis performance

➔ **Deep Generative Models**



Generative Adversarial Networks – GANs



Generating Tabular Data in the Medical Field

GANs

- Impressive results in generating images
- Challenging performance on **tabular data** especially in medical domain with **limited data**

GAN variants

- generate tabular data
- consider limited amount of data (small subsets of real data)

Breast Cancer Wisconsin Data

data patients with breast tumors

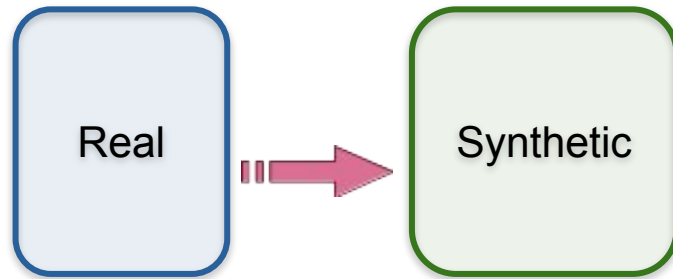
Primary task: Binary Classification (cancerous / non-cancerous tumors)

	Radius	Texture	Perimeter	Area	Smoothness	Compactness
0	17.99	10.38	122.80	1001.0	0.11840	0.27760
1	20.57	17.77	132.90	1326.0	0.08474	0.07864
2	19.69	21.25	130.00	1203.0	0.10960	0.15990
3	11.42	20.38	77.58	386.1	0.14250	0.28390
4	20.29	14.34	135.10	1297.0	0.10030	0.13280

Wolberg, W.; Street, W.; Mangasarian, O. Breast Cancer Wisconsin (Diagnostic); UCI Machine Learning Repository. 1995.

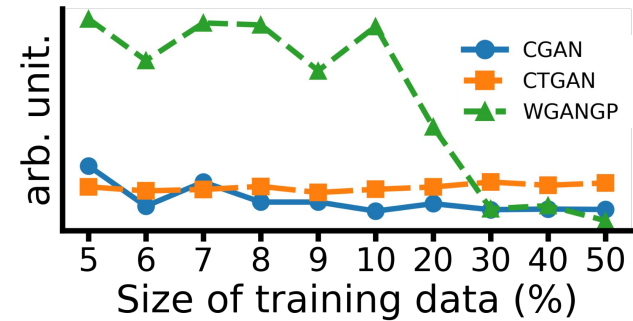
Quality of Synthetic Data

- Conventional methods
characteristics of real data captured by synthetic data

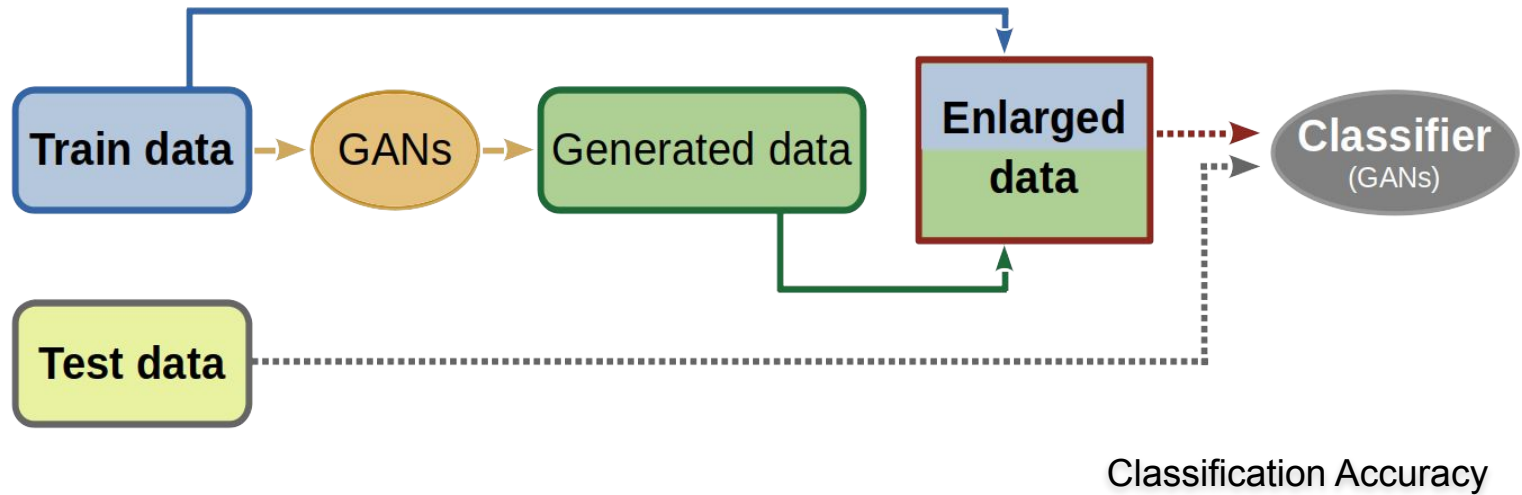


- Target application: Binary Classification

Pairwise Correlation Distance

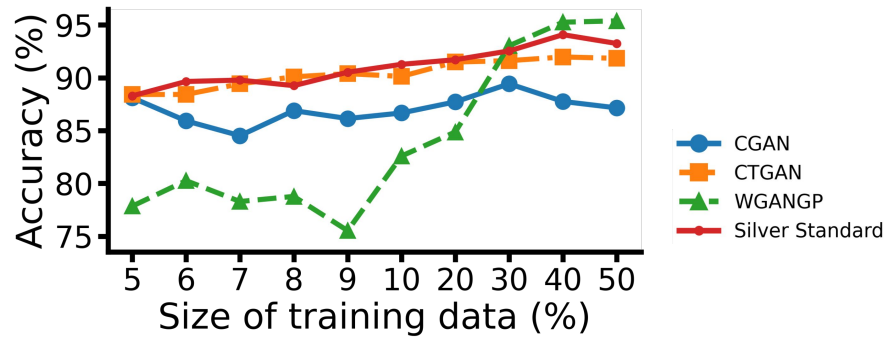


Evaluation Framework



Abedi, M.; Hempel, L.; Sadeghi, S.; Kirsten, T. **GAN-Based Approaches for Generating Structured Data in the Medical Domain**. Appl. Sci. 2022, 12, 7075.

Accuracy of Classifiers using data from GANs



Improved accuracy by including synthetic data

- ❑ compared to Silver Standard (with only real data)
- ❑ even with small amount of available data

Abedi, M.; Hempel, L.; Sadeghi, S.; Kirsten, T. **GAN-Based Approaches for Generating Structured Data in the Medical Domain**. Appl. Sci. 2022, 12, 7075.

Privacy-preserving Diagnostic Support for Leukodystrophy Patients

Navid Shekarchizadeh



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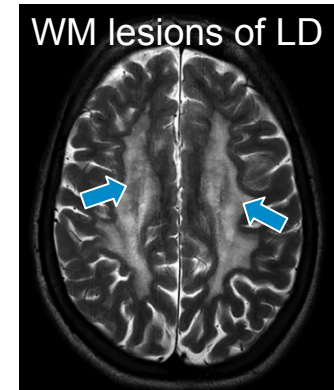
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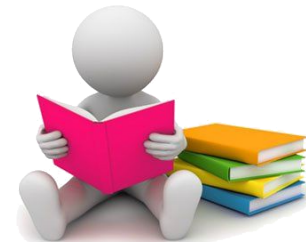
Close collaboration with local
clinicians of the MZEB ❤️

LEUKODYSTROPHIES: Introduction

- Leukodystrophies (LD):
 - A group of genetic disorders
 - Neurodegenerative
 - Rare
 - Fatal
 - Differential diagnosis: Multiple Sclerosis (MS)
- **Goal:** Developing a diagnostic advisory system
 - Reusing the data of medical centers
 - Federated learning of artificial intelligence models
 - Preserving patients' privacy




Resende, L., et al. Radiographics 39. (2019)



Best Poster Award: BMG, Nov. 2022

CLASSIFICATION APPROACH FOR POTENTIAL LEUKODYSTROPHY PATIENTS

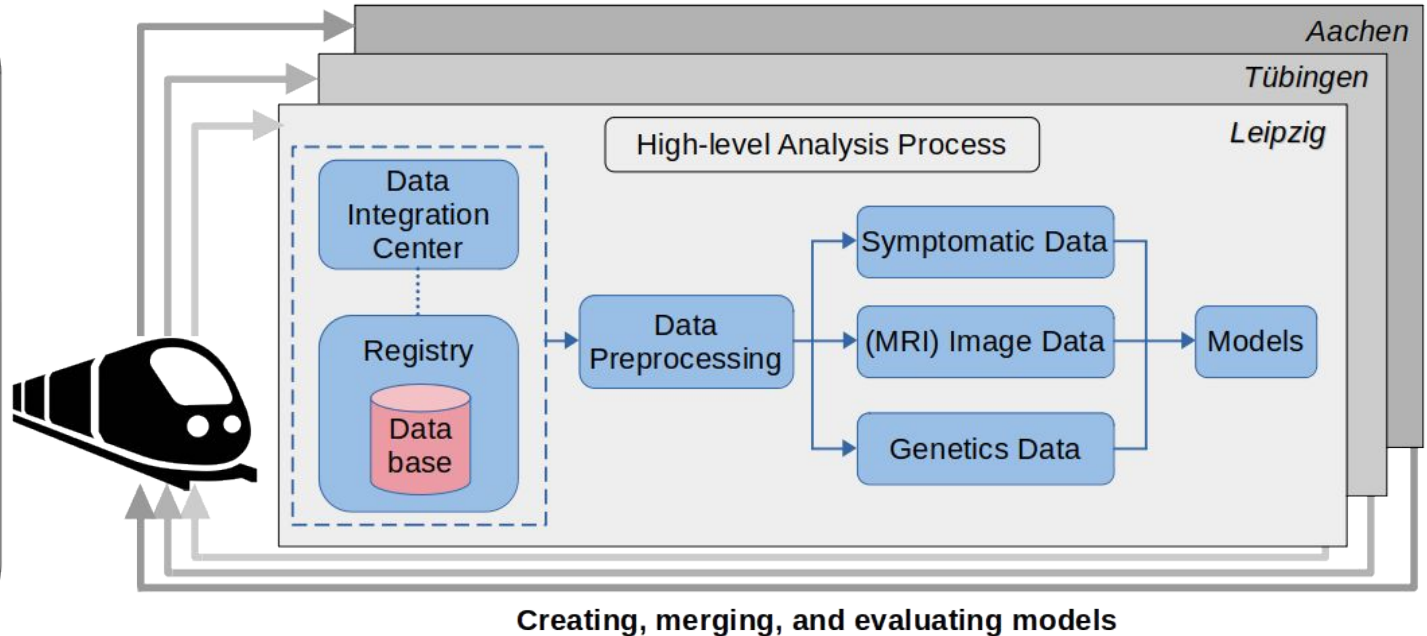


PERSONAL
HEALTH TRAIN

PADME

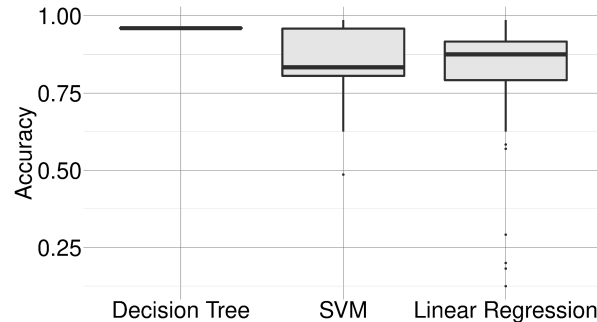
<https://padme-analytics.de>

- ✓ Secure onsite data analysis and result shipping
- ✓ Community-based analysis code review
- ✓ Free of programming languages and data types



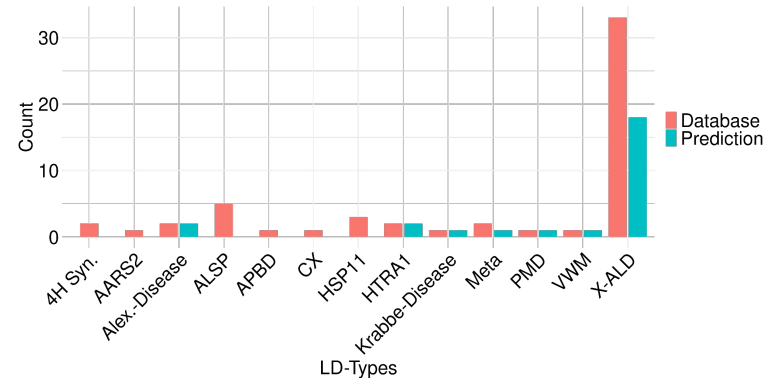
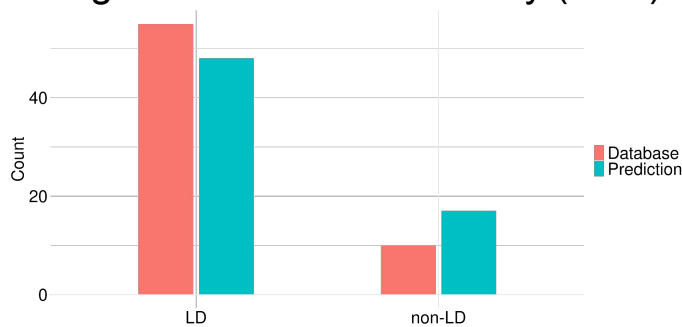
DIFFERENT DATA TYPES

- Symptomatic data
 - Questionnaires and input forms
 - With HPO annotations in REDCap
 - Enlarged data volume by generating synthetic data
- Results
 - Stable accuracy for LD / non-LD classification



DIFFERENT DATA TYPES

- Genetics data
 - Rule-based approach using PhenoMan system (TOP JRG)
 - Reuse of publicly available genetic annotations (e.g., ClinVar)
- Results
 - High classification accuracy (91%)



Publication:

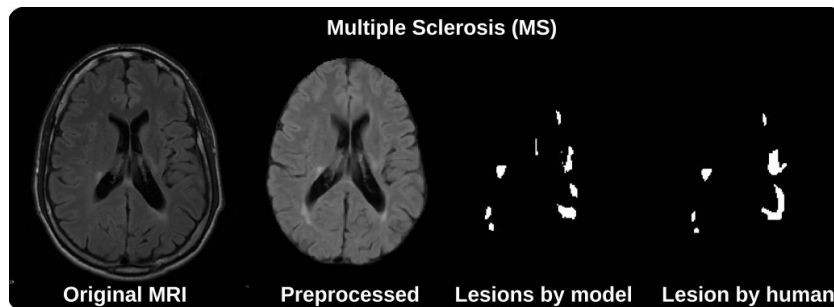
Uciteli A, Beger C, Wagner J, Kiel A, Meineke FA, Stäubert S, Löbe M, Hänsel R, Schuster J, Kirsten T, Herre H. **Ontological Modelling and Execution of Phenotypic Queries in the Leipzig Health Atlas.** Stud Health Technol Inform. 2021 May 24;278:66-74. doi: 10.3233/SHTI210052. PMID: 34042877.

DIFFERENT DATA TYPES

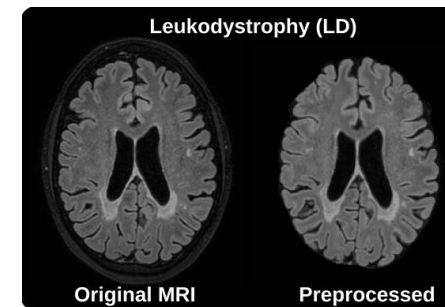
- MRI data
 - Exported and managed by a XNAT system
 - Completed pre-processing pipeline: de-identification, skull stripping, and registration
 - Implemented a deep learning algorithm for MS lesion segmentation (similarity to LD)
 - Challenges:
 - Harmonizing the images from different MRI scanners and configurations
 - Applying the lesion segmentation algorithm to LD (rare disease)

- Results

– Promising lesion segmentation of MS



– Preprocessing LD images



Predicting the Length of Stay in a Hospital

Lars Hempel



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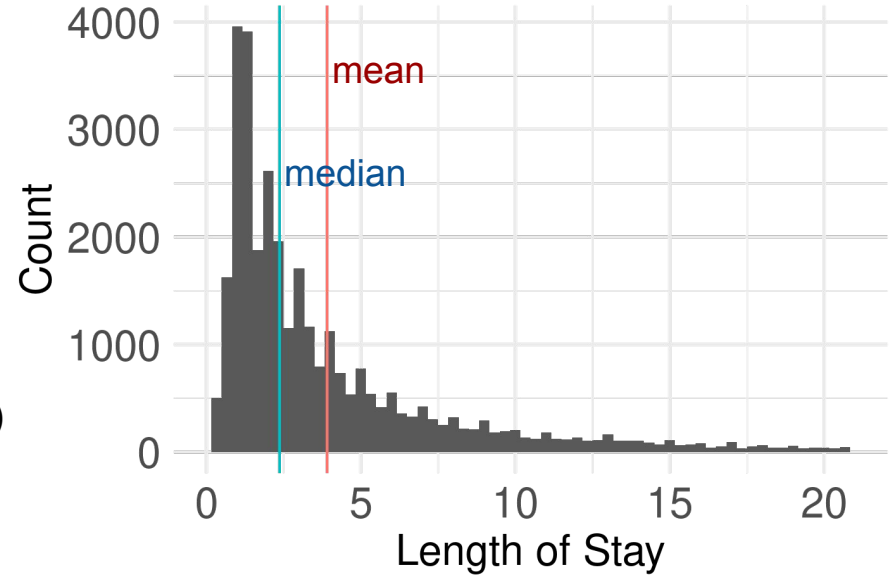


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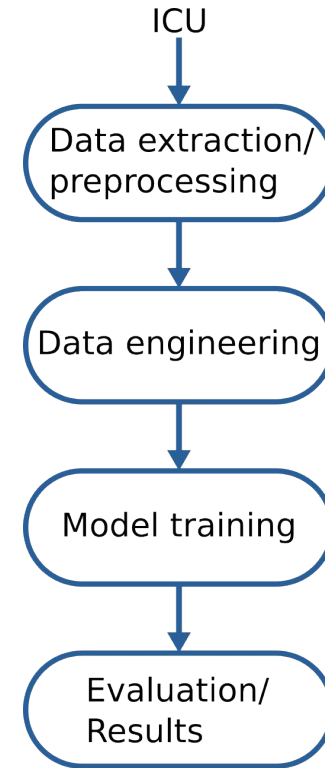
Introduction

- Resource management
 - Indicator for hospital efficiency
 - Reducing costs
- Data: MIMIC IV
 - Demographics (e.g. Age, Sex)
 - Examination (e.g. height, weight)
 - Lab (e.g. Glucose/Potassium Concentration)



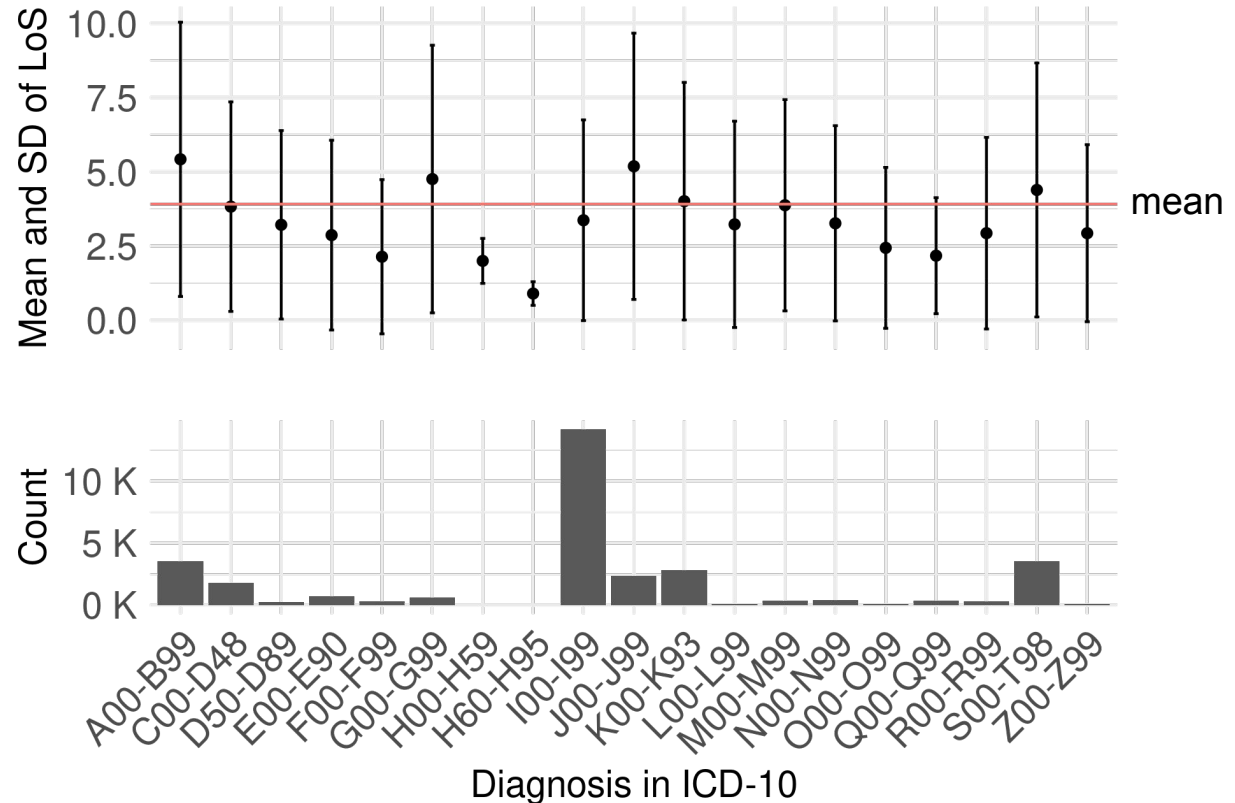
Preprocessing & Methods

- Extract from the database all ICU patients
- Remove missing data
- Filter on:
 - Only adults (age over 18)
 - Weight smaller than 30 Kg
 - Height bigger than 250 cm
 - Exclude patients who died within the stay
- Methods: mean, median, linear Regression, Support Vector Machine and Random Forest
- Split Train/Test (80/20) 10 times
- Evaluation metrics: RMSD, MAD and R^2



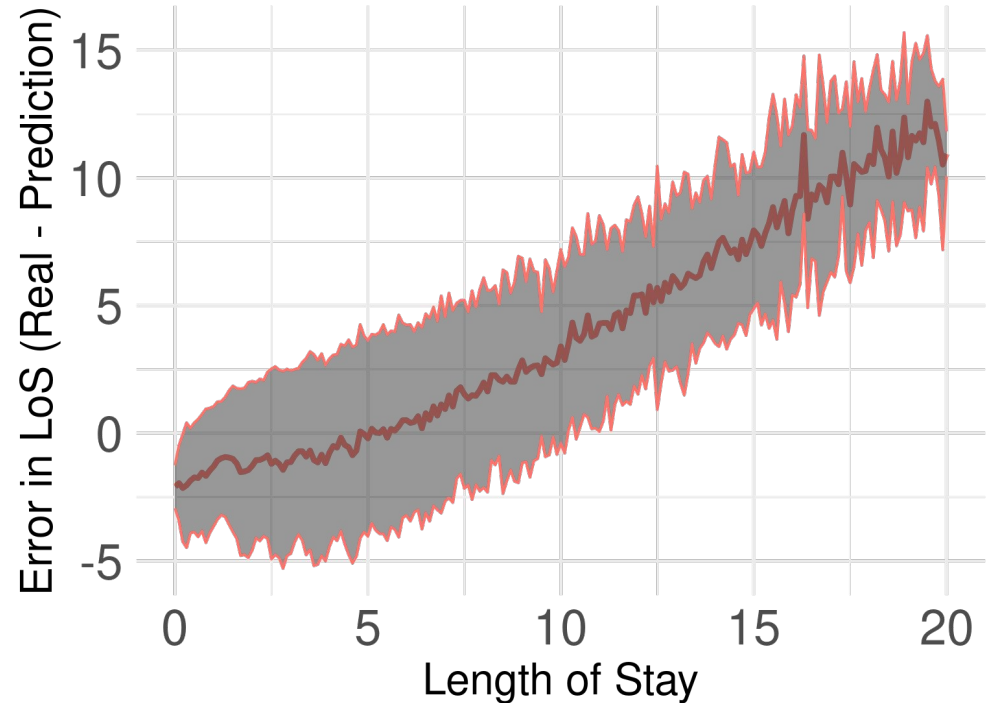
Data exploration Diagnosis

- #stay: 31916
- #features: 20
- mean: 3.91
- median: 2.37
- ICD-10 Coding System
- 14183 in I00 to I99
-> Cardiovascular Diseases



Results

- Best model: Random Forest
- Problems with larger Length of Stay in the Database
- RMSD: 2.88
- Future work:
 - Mixed models for long and short stays
 - Transfer knowledge to hospital data



Early Detection of Sepsis: The AMPEL Project

Daniel Steinbach



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Early Detection of Sepsis

Dr. med. Daniel Steinbach





Leipzig



Grimma



Wurzen





Belegung mit 23 Patienten

Zimmer	Bett	Gsp.	Patient/Geschl./Alter	GebDatum	AMPEL	R	Studie	Abw.	PP
F4092	F04X1014	☒							
F4090	F04X1010	☒							
F4094	F04X1017				○○○				
F4084	F04X1002				○○○				
F4097	F04X1021				○○○		☒		
F4092	F04X1013					⚠			
F4091	F04X1012				●○○	⚠			
F4098	F04X1024				○○○				
F4084	F04X1001				○○○				
F4086	F04X1003					⚠			
F4095	F04X1019				○○○				
F4088	F04X1005				○○○		⚠		
F4098	F04X1023								
F4095	F04X1020				○●○				

Automatisch generierter Hinweis aus dem AMPEL-Meldesystem

Es handelt sich um ein Forschungssystem zur Unterstützung der Patientensicherheit. Auf Grund des Forschungscharakters und der Randomisierung kann die Verfügbarkeit und Validität der Alarme nicht zugesichert werden. Es darf nicht allein aus der Alarmierung eine medizinische Konsequenz abgeleitet werden. Die zugrundeliegenden Befunde müssen geprüft werden.

Website: www.ampel.care | laboratoriumsmedizinischer Dienst: 22221

Name	Geschlecht	Alter	Ausgabe am
Einrichtung 1	Fall	Patient	Uhrzeit



- ⚠ AM-ANÄM-L** ▼
- AKIN ▼
- HYPERKALZIÄMIE ▼
- HYPOKALIÄMIE ▼
- LAKTAT ▼
- HYPONATRIÄMIE ▼
- REFEEDING SYNDROM ▼

Automatisch generierter Hinweis aus dem AMPEL-Meldesystem

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Website: www.ampel.care | laboratoriumsmedizinischer Dienst: 22221

Name	Geschlecht	Alter	Ausgabe am
Einrichtung	Fall	Patient	Uhrzeit



⚠ AM-ANÄM-L

Status	Angelegt am	Angelegt um	Analyt	Ergebnis	Quittiert am	Quittiert um	Aktion
⚠	10.11.2022	09:42:31	Hämoglobin	2,3 mmol/l			Quittieren
✓	09.11.2022	12:42:49	Hämoglobin	2,1 mmol/l			
✓	07.11.2022	10:44:53	Hämoglobin	2,6 mmol/l			
✓	04.11.2022	11:56:47	Hämoglobin	2,4 mmol/l			
✓	02.11.2022	12:14:30	Hämoglobin	2,6 mmol/l			

☑ AKIN

☑ HYPERKALZIÄMIE

Automatisch generierter Hinweis aus dem AMPEL-Meldesystem

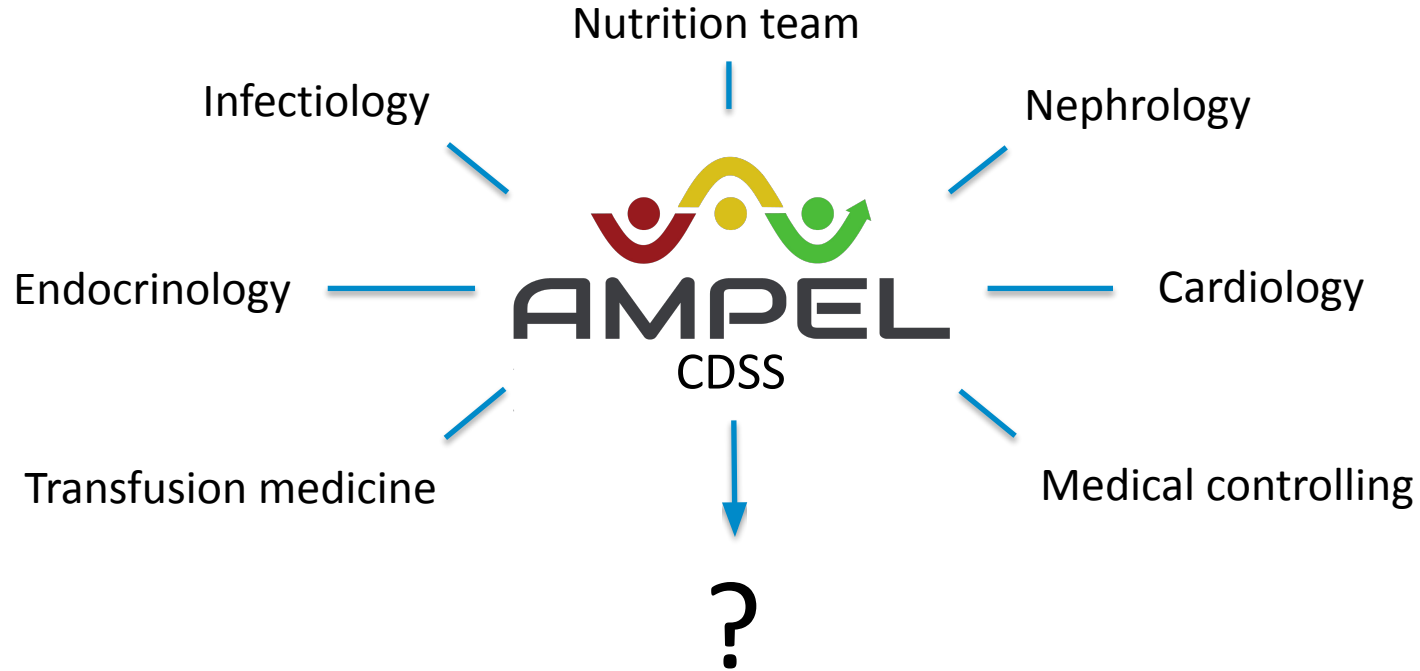
Es handelt sich um ein Forschungssystem zur Unterstützung der Patientensicherheit. Auf Grund des Forschungscharakters und der Randomisierung kann die Verfügbarkeit und Validität der Alarme nicht zugesichert werden. Es darf nicht allein aus der Alarmierung eine medizinische Konsequenz abgeleitet werden. Die zugrundeliegenden Befunde müssen geprüft werden.

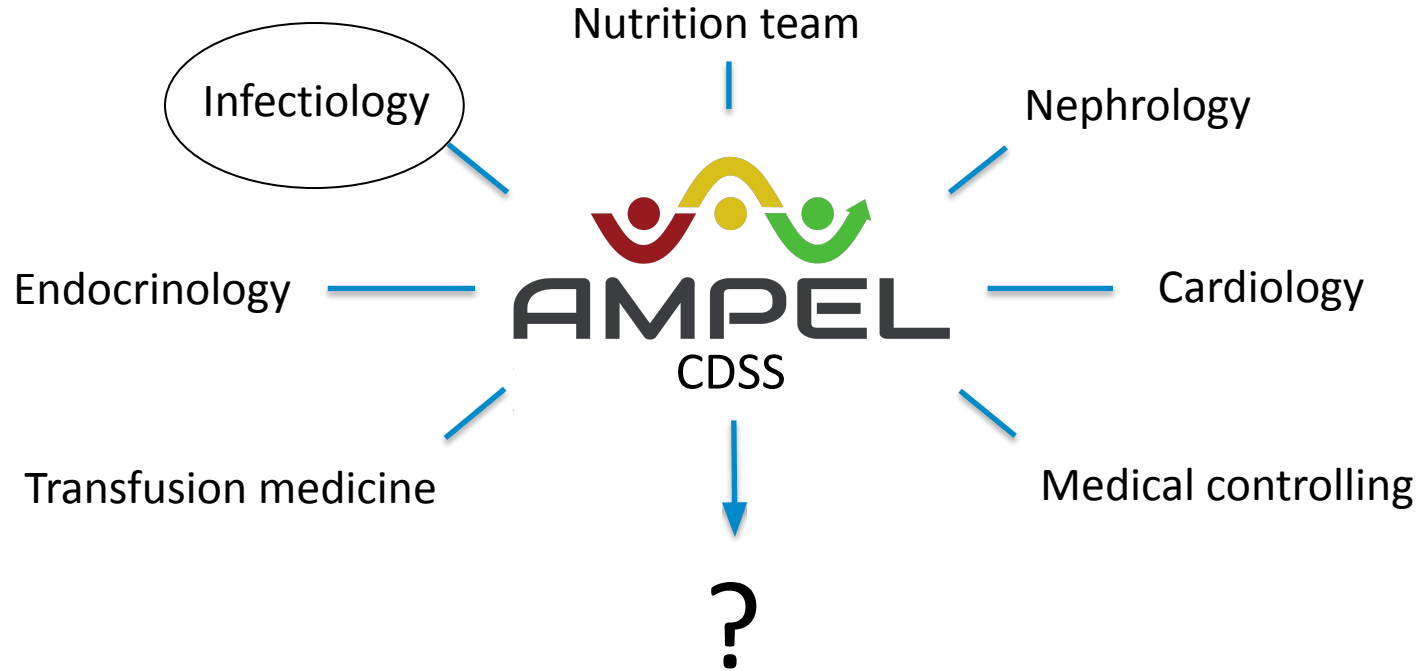
Website: www.ampel.care | laboratoriumsmedizinischer Dienst: 22221

Name	Geschlecht	Alter	Ausgabe am
Einrichtung	Fall	Patient	Uhrzeit
1			



- ⚠ AM-ANÄM-L** ▼
- AKIN ▼
- HYPERKALZIÄMIE ▼
- HYPOKALIÄMIE ▼
- LAKTAT ▼
- HYPONATRIÄMIE ▼
- REFEEDING SYNDROM ▼







Blood Count (BC)

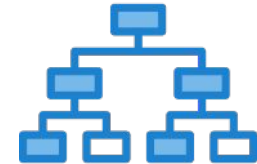


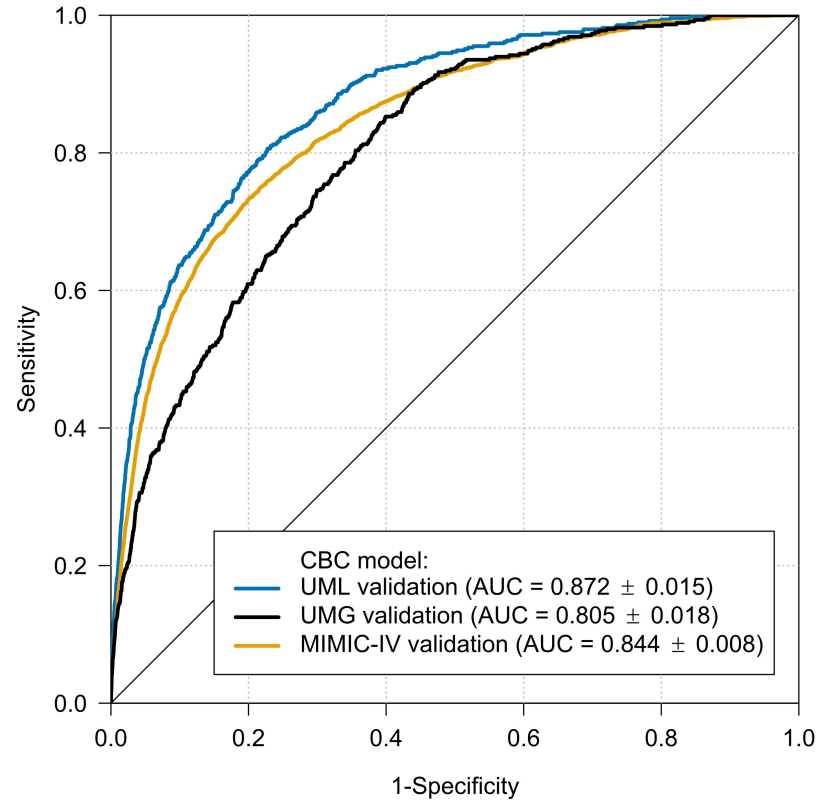
Label

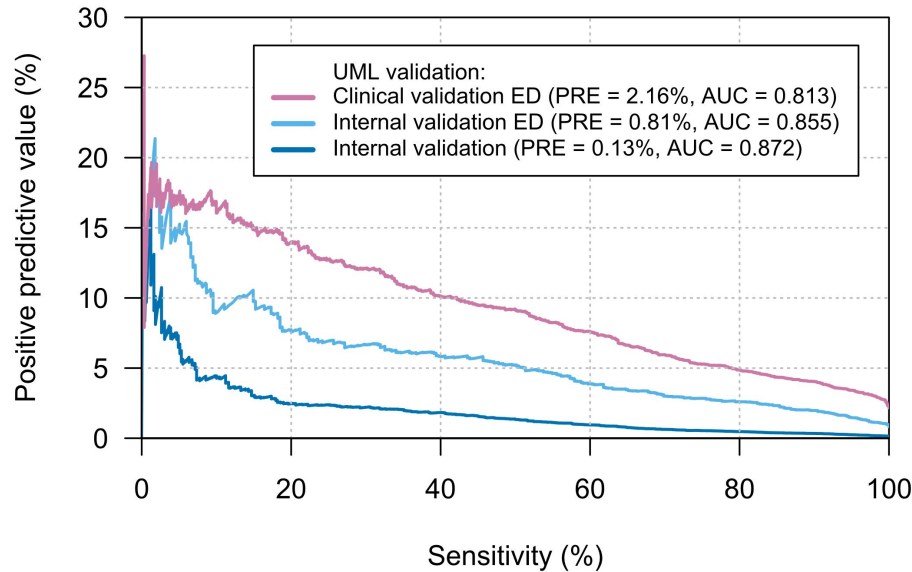
0-6h before ICU
admission with sepsis



Random Forest







medRxiv



BMJ Yale

THE PREPRINT SERVER FOR HEALTH SCIENCES

Applying Machine Learning to Blood Count Data Predicts Sepsis with ICU Admission

Paul C. Ahrens, Daniel Steinbach, Maria Schmidt, Martin Federbusch, Lara Heuft, Christoph Lübbert, Matthias Nauck, Matthias Gründling, Berend Isermann, Sebastian Gibb, Thorsten Kaiser

doi: <https://doi.org/10.1101/2022.10.21.22281348>

Dataset
published
Code
published
R-Packages
published





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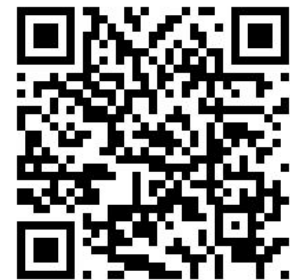
Medizin ist unsere Berufung.

„Come Together“

- John Lennon



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sepsis preprint

THANK YOU!

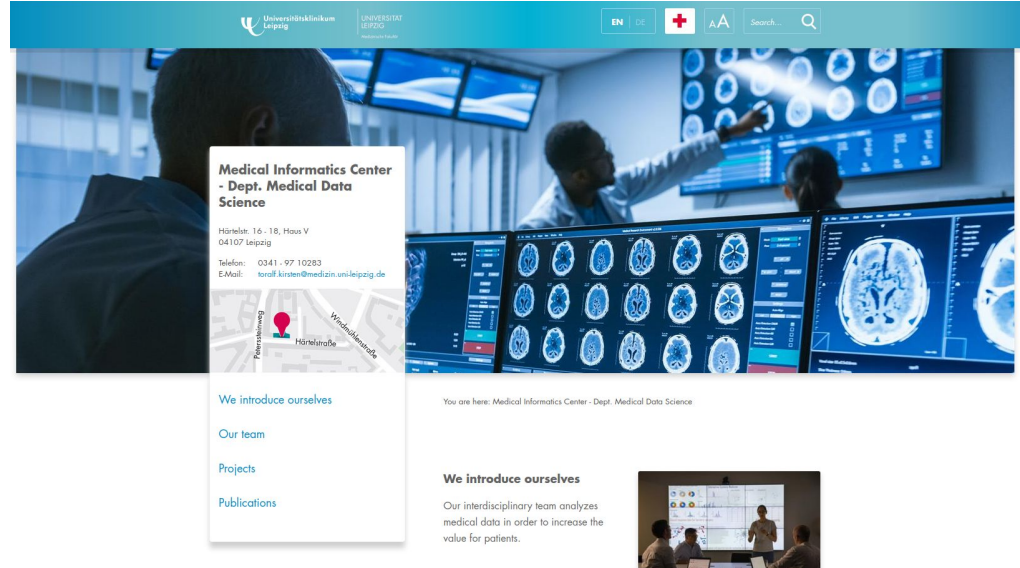
Toralf Kirsten
Medical Data Science

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T +49 341 97-10283

toralf.kirsten@medizin.uni-leipzig.de

<https://www.uniklinikum-leipzig.de/einrichtungen/medical-data-science>

There are refreshments and posters in front of this lecture hall. We are delighted to answer questions.





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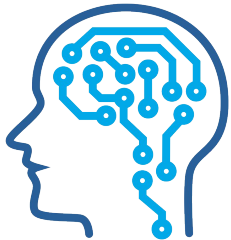


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Posters

Leipzig, 02.12.2022



MDS

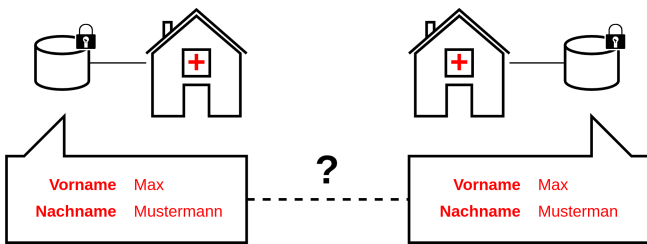
Medical Data Science

Privatsphärenschützende Datenverknüpfung in verteilten Analysen mit dem PHT

Maximilian Jug^{1,2,3}, Sascha Welten⁴, Yongli Mou⁴, Yeliz Ucer Yediel⁵, Oya Beyan⁵, Ulrich Sax⁶, Toralf Kirsten^{1,2,3}

Affiliations: 1) Abteilung Medical Data Science, Medizininformatikzentrum Leipzig 2) Institut für medizinische Informatik, Statistik und Epidemiologie, Universität Leipzig 3) Fakultät Angewandte Computer- und Biowissenschaften, Hochschule Mittweida 4) Lehrstuhl Informatik 5, RWTH Aachen 5) Abteilung Data Science und Künstliche Intelligenz, Fraunhofer FIT 6) Institut für Medizinische Informatik, Universitätsmedizin Göttingen

Hintergrund



- Verteilte Datensätze und hohe Datenschutzanforderungen
- Personal Health Train (PHT) als Plattform für verteilte Analysen[1]
- Dubletten können statistische Ergebnisse verzerren
- Record Linkage (RL) zur Verbindung von Datenquellen
- Ziel: Integration von RL-Methoden (PPRL) in den PHT

Methoden

- Erweiterung der PHT-Implementierung PADME[2]
- PPRL mit Bloomfilter für Datenverknüpfung[3]
- Durchführung in zwei Zugausführungen im PHT

Stationsdienste

Encoder

- Vorverarbeitung von IDATs
- Maskierung von IDATs

Resolver

- Auflösung von Pseudonymen
- Übermittlung von Bitvektoren

Zentrale Dienste (TTP)

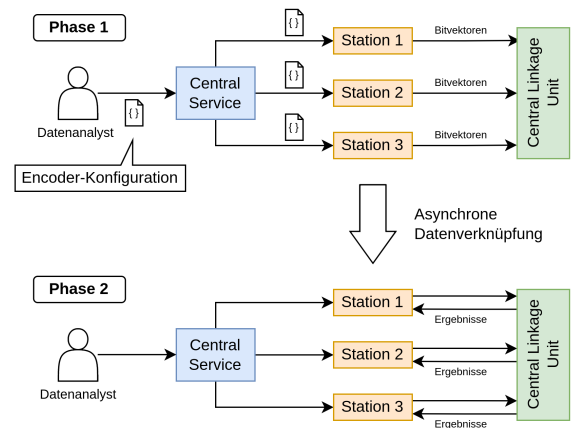
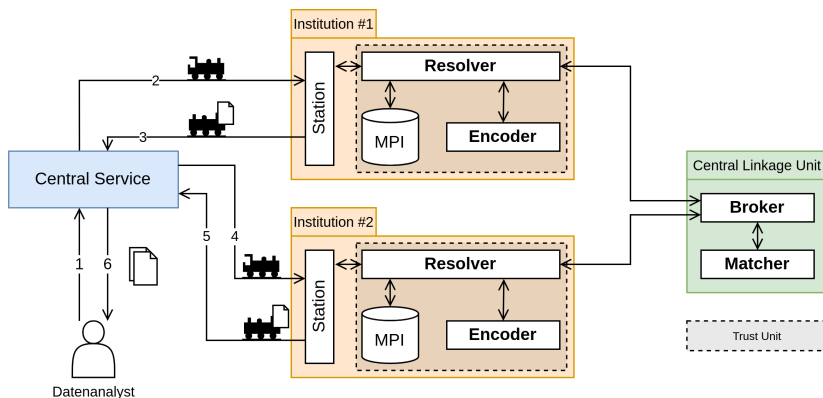
Broker

- Annahme von Bitvektoren
- Verwaltung von Match-Sessions
- Bereitstellung von Ergebnissen

Matcher

- Abgleich von Bitvektoren

Architektur, Komponenten und Prozess



Ergebnisse

- Validierung des Verfahrens an zwei Datensätzen
- CORD „Private Set Intersection“ → **100 % max. F1-Score**
- Synthetischer Datensatz → **99,7 % max. F1-Score**
→ IDATs aus öffentlichen Datenquellen
→ Typografische Fehler für Fehlertoleranz des Verfahrens

Diskussion

- Kein Unterschied zur zentralisierten Datenverknüpfung
- Anwendung von Strategien zur Behandlung von Dubletten → **Entity Consolidation und Duplicate Elimination**
- Sichere Verknüpfung durch Härtung des PPRL-Verfahrens → **Keine Rückschlüsse auf IDATs möglich**

Referenzen: [1] O. Beyan u. a., „Distributed Analytics on Sensitive Medical Data: The Personal Health Train“, Data Intelligence, Bd. 2, Nr. 1–2, S. 96–107, Jan. 2020, doi: 10.1162/dint_a_00032. [2] S. Welten u. a., „A Privacy-Preserving Distributed Analytics Platform for Health Care Data“, Methods Inf Med, Jan. 2022, doi: 10.1055/s-0041-1740564. [3] R. Schnell, T. Bachteler, und J. Reiher, „Privacy-preserving record linkage using Bloom filters“, BMC Med Inform Decis Mak, Bd. 9, Nr. 1, S. 41, Aug. 2009, doi: 10.1186/1472-6947-9-41.

Generating Structured Data in the Medical Domain using GANs

Sina Sadeghi, Lars Hempel, Masoud Abedi, Toralf Kirsten

Department for Medical Data Science, Leipzig University Medical Center
 Institute for Medical Informatics, Statistics and Epidemiology, Leipzig University
 Faculty Applied Computer and Bio Sciences, Mittweida University of Applied Sciences



Introduction

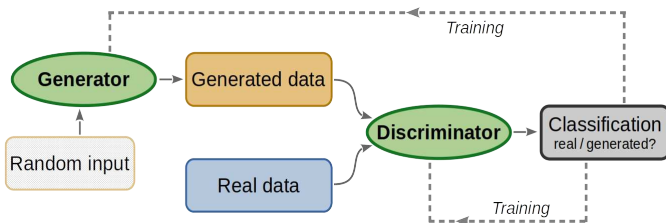
Medical data scarcity:

- ❖ Patient privacy (data protection regulations)
- ❖ Rare diseases with limited available data

Synthetic data generation:

- ❖ Artificial data resembling real data
 - not associated with real patients
 - improve data analysis performance

GANs - Generative Adversarial Networks



- ❖ Impressive results in generating images
- ❖ Challenging performance on tabular data
 - especially in medical domain with limited data

Methods

GAN variants for generating tabular data:

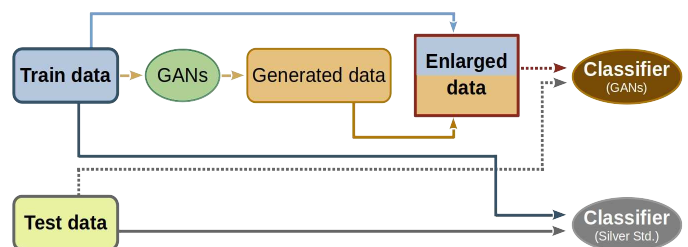
- ❖ CGAN - Conditional GANs
- ❖ CTGAN - Conditional Tabular GANs
- ❖ WGANGP - Wasserstein GANs with Gradient Penalty

BCW - Breast Cancer Wisconsin dataset

- ❖ Differentiating into small subsets
 - effect of size of training data on model accuracy
- ❖ Regular extension sampling
 - impact of newly available on model performance

Evaluation framework

Extended data consisting of real and synthetic data



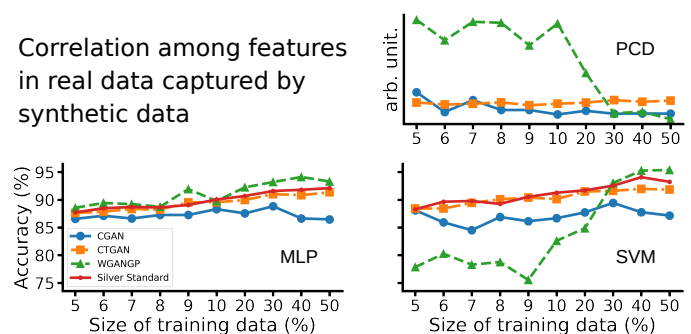
Binary (malignant/benign) classifiers:

- ❖ SVM - Support Vector Machine
- ❖ MLP - Multi-Layer Perceptron

Results

PCD - Pairwise Correlation Difference

Correlation among features in real data captured by synthetic data



Improved accuracy by including synthetic data

- compared to Silver Standard (with only real data)
- even with small amount of available data

Other ratios for real and synthetic data in the extended dataset are be considered

Reference

Abedi, M.; Hempel, L.; Sadeghi, S.; Kirsten, T. GAN-Based Approaches for Generating Structured Data in the Medical Domain. Appl. Sci. 2022, 12, 7075.

An Image Processing Pipeline to Detect Potential Leukodystrophy Patients

Masoud Abedi^{1,2,3} Navid Shekarchizadeh^{1,2} Sina Sadeghi^{1,2} Lars Hempel^{1,2,3}
 Christa-Caroline Bergner⁴ Julia Lier⁴ Wolfgang Köhler⁴ Toralf Kirsten^{1,2,3}
 on behalf of the LEUKO-Expert Project Team

- 1) Department of Medical Data Science, Leipzig University Medical Center
- 2) Institute for Medical Informatics, Statistics and Epidemiology, Leipzig University
- 3) Faculty of Applied Computer and Bio Sciences, Mittweida University of Applied Sciences
- 4) Department of Neurology, University of Leipzig Medical Center



Introduction & Methods

- Leukodystrophies (LD):
 - A family of neurodegenerative diseases
 - ✓ Image-based diagnosis is crucial
 - Rare (1 in 40,000 people worldwide)
 - ✓ Synthetic data generation is necessary
 - MRI data available in 3 hospitals in Germany
 - ✓ Distributed & privacy-preserving data analysis
- Multi-site image processing pipeline:
 - Diagnostic support, education, and training
 - Personal Health Train (PHT) infrastructure [1]
 - White matter lesion segmentation modeling
 - ✓ Multiple Sclerosis (MS) is very similar to LD
 - ✓ Annotated data from ISBI challenge 2015 [2]
 - ✓ Preprocessing
 - ✓ Deep convolutional neural network

Results

- Promising performance of the model on MS
- Preprocessing MRI of LD (data not annotated)

Multiple Sclerosis (MS)

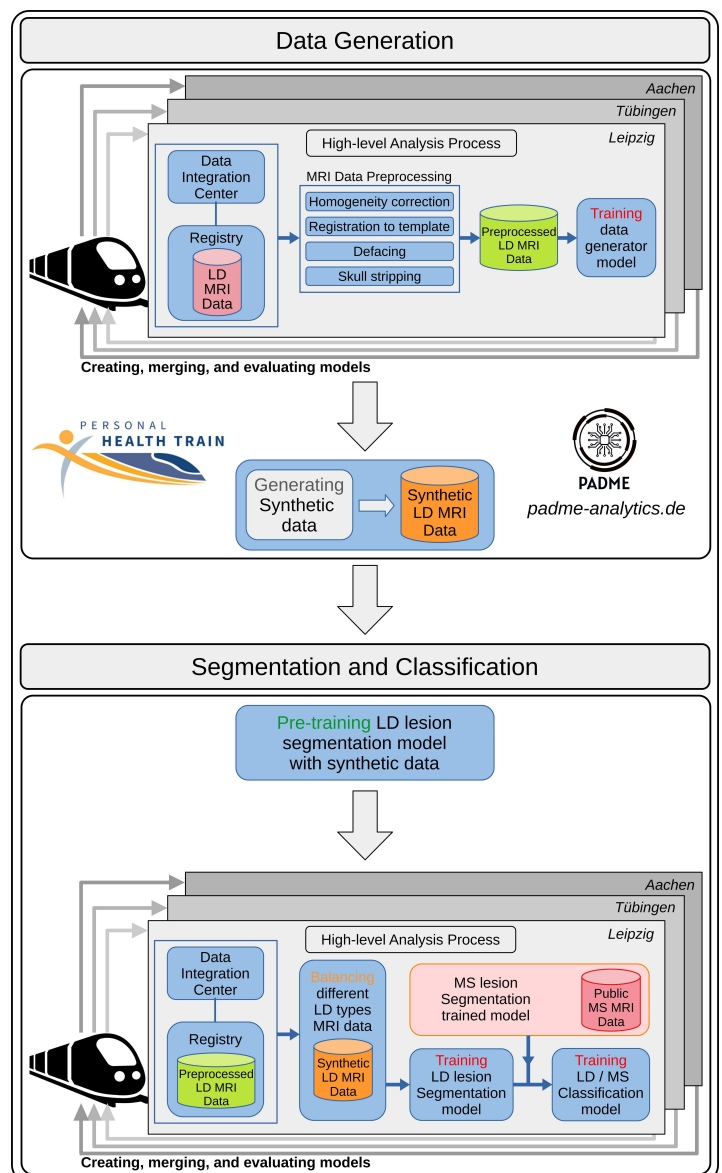
Leukodystrophy (LD)

Metric	Value
Lesion false positive rate	0.29
Dice coefficient	0.67
Positive predictive value	0.71
Volume difference	0.32

Future work

- Transforming the model from MS to LD
- LD MRI data harmonization

Multi-Site Image Processing Pipeline



Vorhersage der Verweildauer auf einer ITS

Lars Hempel^{1,3}, Sven Bercker², Samira Zeynalova³, Sina Sadeghi^{1,3}, Ulrike Klotz^{1,3,4},
Toralf Kirsten^{1,3,4}

- 1) Abteilung Medical Data Science, Medizininformatikzentrum Leipzig
- 2) Klinik und Poliklinik für Anästhesiologie und Intensivtherapie, Universitätsklinikum Leipzig
- 3) Institut für medizinische Informatik, Statistik und Epidemiologie, Universität Leipzig
- 4) Fakultät Angewandte Computer- und Biowissenschaften, Hochschule Mittweida

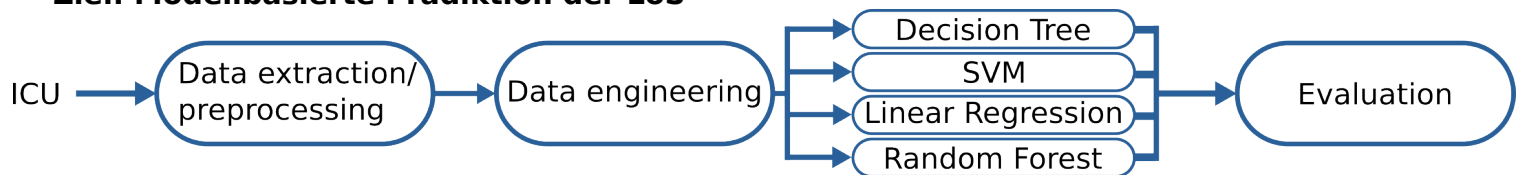


Motivation

- Ressourcenmanagement als Indikator für Effizienz und Kostenverwaltung
- Aufenthaltsdauer eines Patienten (LoS) für die Bewertung des Managements und Pflege
- Aufenthaltsdauer abhängig von Patienten, dessen Symptomatik und Kondition
- **Ziel: Modellbasierte Prädiktion der LoS**

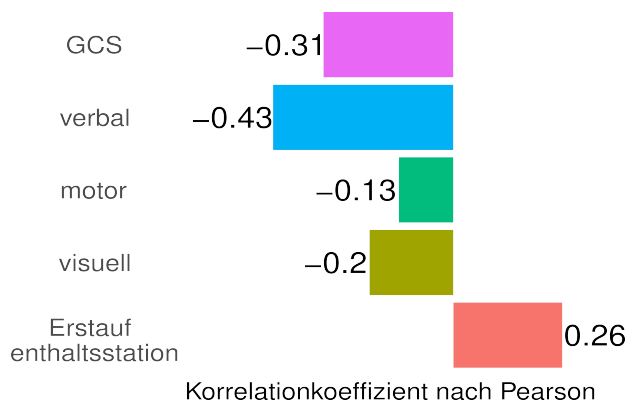
Methoden

- Datenquelle: ITS-Daten aus MIMIC IV
- Datenqualität: Konservative Kuratation auf Basis von Literatur und med. Experten
- Methoden zur Prädiktion: Decision Tree, SVM, Linear Regression und Random Forest
- Evaluationsmetriken: RMSD, MAD, MAPE, R²



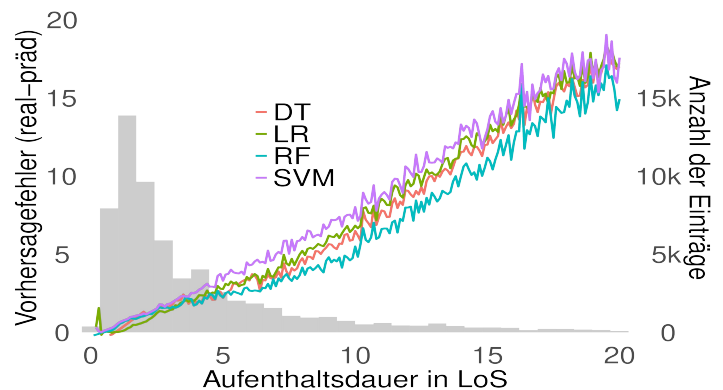
Ergebnisse

- Beste Modellprädiktion: **Random Forest**
- RMSD: 2,88 und R²: 0,41
- Kohortengröße: 24 148, genutzte Parameter: 20
- Wichtigste Einflussgrößen: Glasgow Coma Scale (GCS) und Erstaufenthaltsstation
- Viele Laborparameter mit geringem Einfluss
- Größerer Prädiktionsfehler bei steigenden LoS



Diskussion

- Ansatz: Prädiktion auf der Basis der Erstanamnese mit akzeptablen Ergebnissen
- Prädiktionfehler bei großen LoS auf Grund von kleinen Fallzahlen und ggfs. ungeplanten Ergebnissen
- Zukünftige Ausrichtung: Kontinuierliche Adaption der LoS Prädiktion unter Beachtung longitudinaler Diagnostik



Referenzen

- 1) Bacchi, S. et al. (2022), Machine learning in the prediction of medical inpatient length of stay. Intern Med J, 52: 176-185. <https://doi.org/10.1111/imj.14962>
- 2) Rocheteau, E. et al. (2021) Temporal pointwise convolutional networks for length of stay prediction in the intensive care unit. In Proceedings of the Conference on Health, Inference, and Learning (CHIL '21). Association for Computing Machinery, New York, NY, USA, 58-68. <https://doi.org/10.1145/3450439.3451860>



Retinal layer thickness as a biomarker for monitoring the progression of diseases of the central nervous system –

Medical Data Science using OCT and MRI imaging

S.Sadeghi¹, T. Kirsten^{1,2}, S. Klumpe³, M. Eulitz³, A. Dillenseger³, C. Beste⁴, F. Ziemssen⁵, T. Ziemssen³, F.G.Rauscher^{1,2}

¹⁾ Medical Data Science, Universitätsklinikum Leipzig, Leipzig, ²⁾ Institut für Medizinische Informatik, Statistik und Epidemiologie, Universität Leipzig, Leipzig, ³⁾ Zentrum für Klinische Neurowissenschaften, Klinik und Poliklinik für Neurologie, Technische Universität Dresden, Dresden, ⁴⁾ Klinik und Poliklinik für Kinder- und Jugendpsychiatrie und Jugendpsychotherapie, Technische Universität Dresden, Dresden, ⁵⁾ Augenklinik, Universitätsklinikum Leipzig, Leipzig

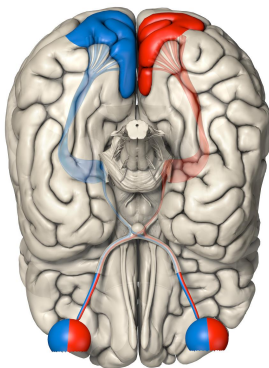
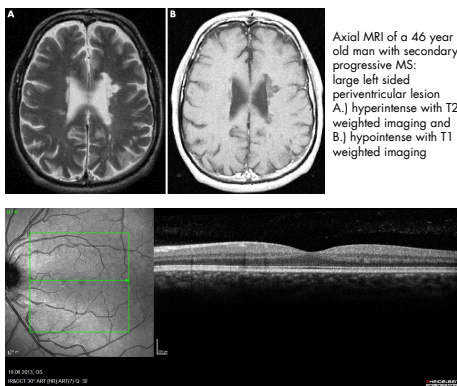
Background

The eye is a window to the brain. There is promising evidence of correlations between retinal layer thickness from optical coherence tomography (OCT) and magnetic resonance imaging (MRI) markers.

→ This results in the potential for science-based clinical application of OCT and MRI in combination for diagnosis and follow-up of various diseases of the central nervous system.

■ In the present study, retinal structures (measured with OCT) are compared with intracerebral structures and pathologies from available MRI examinations. In addition, their correlations to cognitive functions will also be investigated.

■ For this purpose, existing OCT, MRI and data from neuropsychological and cognitive examinations (partly under EEG) of patients of the ZKN and test persons of the ZKN and KJP will be used (retrospective study design).



Methods

complete dataset: 316,681 clinical images from existing documentation of clinical routine for 2,662 individuals (including data and information collected during clinical routine in connection with the OCT and/or MRI examinations)

removal of duplicates

study population: 85,167 clinical images for 2,659 patients of the ZKN as well as for subjects who received OCT and/or MRI examinations at the ZKN and KJP of the University Hospital Carl Gustav Carus Dresden, Germany

exclusion of 4,293 records of 1,047 patients with non-standardised image settings (excluded scans belong to 9 patients who additionally also possess all standardised scans (=records not lost); or to 1,032 patients with only some of the required imaging; and further to 6 patients not included in the standardised sample).

analysis subset: 64,925 clinical images for 1,659 patients imported into the analysis pipeline

■ Cross-sectional and longitudinal analyses for three key OCT scan types of the Glaucoma Module by Spectralis spectral-domain Optical Coherence Tomography (SD-OCT), Heidelberg Engineering, Heidelberg, Germany

■ Scan settings were:

Macular volume scan with horizontal scan location [PPoleH[APS]] in parallel to the disk-fovea-axis. 61 B-scans with a distance of 120µm between scans inside a 30° by 25° area. High-speed mode with 768 x 496 pixel.

Macular volume scan with vertical scan location [PPoleV[APS]] perpendicular to the disk-fovea-axis. 19 B-scans with a distance of 240µm between scans inside a 30° by 15° area. High-resolution mode with 1536 x 496 pixel.

Circumpapillary ring scan around the optic nerve head [ONH-RC[APS]] comprising of three concentric ring scans with 3.5mm, 4.1mm, and 4.7mm diameter (accompanied by 24 line scans). High-speed mode with 768 x 496 pixel

→ These three scan patterns were matched by regular volume and circular scans of the same resolution settings to increase the number of analyzed patients, when possible.

Results

Standardised SD-OCT imaging at the Multiple Sclerosis Centre Dresden, Centre of Clinical Neuroscience, Department of Neurology, Carl Gustav Carus Faculty of Medicine, Technical University Dresden, Germany was analyzed for fixed scan-settings.

■ Horizontal volume scans with 61 B-Scans, vertical volume scans with 19 B-Scans and one version of circumpapillary Retinal Nerve Fibre Layer scan (either ONH-RC[APS] or one circular B-scan, sometimes combined with 24 radial scans) were present in combination in 1,659 patients for 1,640 right eyes and 1,600 left eyes.

■ Of those, 1,379 (right eye) and 1,347 (left eye) presented with the ideal imaging by the Glaucoma Module (PPoleH[APS], PPoleV[APS] and ONH-RC[APS]). For 422 right eyes and 408 left eyes comparable scan combinations were obtained.

■ In total, 64,925 records of a final sample of 1,659 patients were imported into the analysis pipeline. This clinical dataset consists of routine visits of 1,630 patients (60,233 records) as well as 53 specific study patients with 4,702 records (7% of records).

Image Type	B-Scans
Set 1 OCT Volume Scan	61
Set 2 OCT Circle Scan	1
Set 3 OCT Volume Scan	19
Set 4 OCT Radial Scan	24
Set 5 OCT Radial Circle Scan	27

Exam Date freq	Legend:
Group 1 # <= 2	R1: number of eyes with full set of records for Set 1, Set 3 and Set 5;
Group 2 2 < # <= 4	L1: number of eyes with full set of records for Set 1, Set 3 and Set 5
Group 3 4 < # <= 6	
Group 4 6 < # <= 8	
Group 5 8 < # <= 10	
Group 6 10 < #	

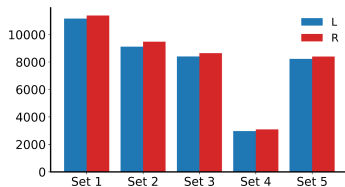


Figure 1 Frequency of desired image settings; Legend: R: right eye; L: left eye

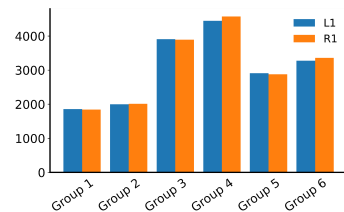


Figure 2 Frequency of examinations with the combination of desired settings: per eye

Discussion

At ZKN, in combination with joint studies by ZKN and KJP, 80,874 images were obtained in the aforementioned volume and circular scan settings, Set 1, Set 3, and Set 5 or comparable (right eye: 40,985; left eye: 39,889). This points to a very high level of standardisation of imaging in this clinical setting. This unprecedented dataset is of high value to the research questions of ZKN and KJP and will serve as the ideal basis for cross-sectional and longitudinal analyses of diseases of the central nervous systems. A substantial number of patients had standardized imaging by the Spectralis Glaucoma Module, for the desired scan settings, 64,925 records of 1,659 patients were available. Inside this sample, 53 specific study IDs were counted, some of which could be the topic of a separate analysis.

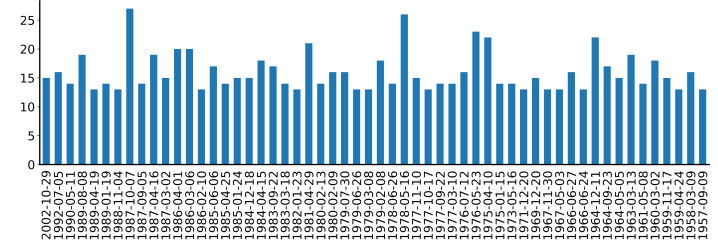


Figure 3 Frequency of exams per patient in Group 6, sorted by birthdate for longitudinal analysis;

Outlook: The 1,659 patients with available 64,925 OCT records will be stratified into clinical research questions to focus on cross-sectional and longitudinal analyses of diseases of the central nervous system, this will include data from MRI and other clinical information. The research will centre on Multiple Sclerosis (MS), with investigations on disease progression, and therapy, with a specific focus on relapsing and remitting MS.

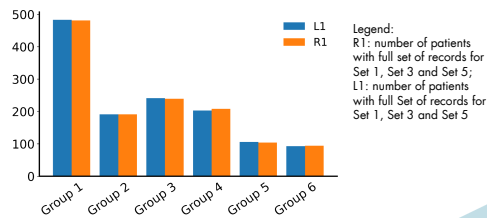
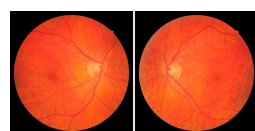


Figure 4 Number of patients with frequency of examinations with the combination of desired settings





Circumpapillary Retinal Nerve Fiber Layer Thickness

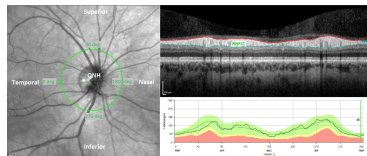
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Motivation

- Early diagnosis of eye disease is of increased importance in an ageing population
- Normative data serve as the backbone for comparison with individual cases
- Such norms need to be accessible directly in clinics to support ophthalmological care
- Ocular optical coherence tomography (OCT) manufacturers aim to mark abnormalities in patient measurements based on their own, typically proprietary norms based on small samples

- Several different visualizations (e.g. sectors and continuous plots) of normative data may help to improve decision-making for clinicians.

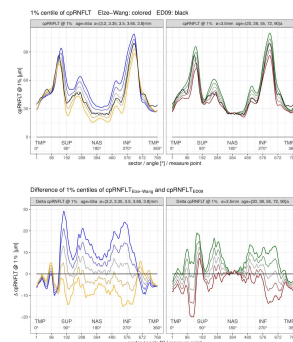
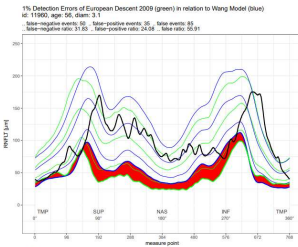


- Circumpapillary Retinal Nerve Fiber Layer Thickness (cpRNFLT) was measured by 768 A-scans on a 360° circle around the optic nerve head of the human eye
- Sectors are commonly used in the clinical context
- Sectors consist of 45° and 90° sections of the scan



Methods

- Normative data calculated from 5637 healthy eyes
- OCT-measurements of the LIFE-Adult-Study population
- Analysing detection errors as a way to judge the improvements of the model by Wang et al. in comparison to the so called "European Descent 2009" (ED09) model used in the OCT device
- Counting false-positive and false-negative detection events assuming the higher accuracy of Elze-Wang's cpRNFLT model

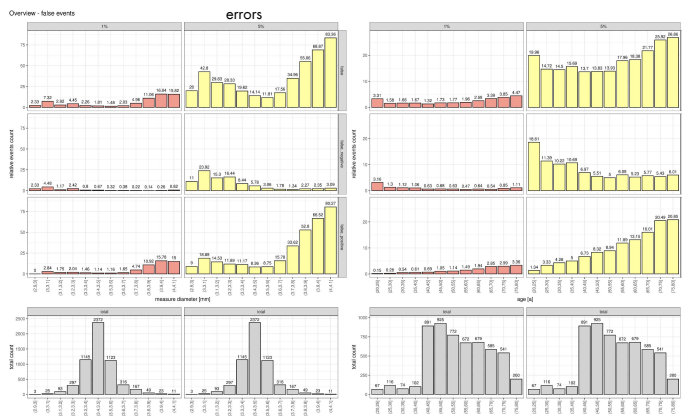
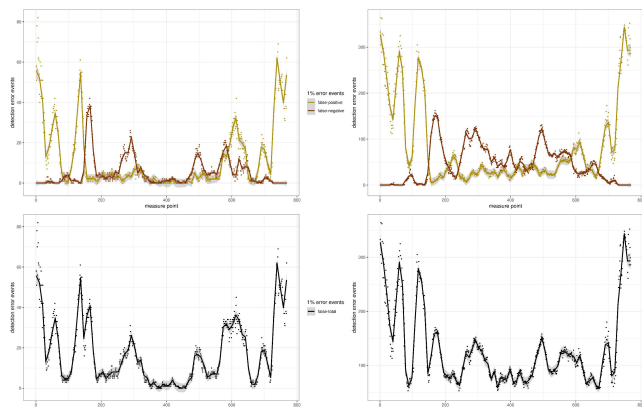


- cpRNFLT 768 measurement points are modelled by a Gaussian distribution
- cpRNFLT-Sectors are modelled by an averaged Gaussian distribution
- cpRNFLT(D) 768 measurement points are modelled by a Box-Cox-T distribution
- cpRNFLT(D)-Sectors-points are modelled by an averaged Box-Cox-T distribution
- Novel cpRNFLT-norms by Wang et al. are modelled as linear dependent on measure diameter and age

Results - Discussion - Outlook

- Influencing factors of the model (e.g. age, measurement diameter) need to be accounted for
- Adjusting for such factors in the model is of direct clinical significance for cpRNFLT as a diagnostic tool
- Dependency of the model on age and diameter are reflected by false-positive and false-negative events
- Thus the RNFLT(D)-Visualizer represents an additional value for clinical decision making in Ophthalmology

- Analysis of relative frequencies of false events for the 1st (red) and the 5th (yellow) percentile with reference to age and measure diameter
- Large diameters and high ages lead to small thicknesses
- Consequently, the rate of false positive errors produced by the built-in ED09 model increases
- Small diameters and low ages are more likely to produce false negative



Successful practical implementation as the RNFLT(D)-Visualizer Shiny-Web-App as part of the Leipzig Health Atlas:

- Opportunity to upload clinical data to enable analysis of this individual level data on the new Wang et al. norms, taking into account age and measurement diameter for the first time
- Importing and loading pdf documents of cpRNFLT measurements and its meta data
- Scanning of OCT print-out of patient measurement plots and read out meta data
- Various presentations and analyses of patient data with device implemented norms and norms from models by Wang et al. via continuous 360° circle plots or sector plots
- Opportunity to download the result graphs and tables in several common formats

References:

Peschel T, Wang M, Krause T, Rauscher FG, Elze T. A Cloud-based Infrastructure for Interactive Analysis of RNFLT Data. In: Heusslein, Vincenz and Buhari, Nina (Hrsg.): EScience@Age 2021. Show Your Research Data. Heidelberg: hdlBOKS; 2022. p.54-63. <https://doi.org/10.11588/hdlBOKS.979.v17718>. **Rauscher FG, Wang M, Franke M, Wilker M, Wehner K, Strass A, Engel C, Thery J, Oberndorf P, Strohbl M, Elze T, Elze T.** Axial function and focal metabolism are major predictors of circumpapillary retinal nerve fiber layer thickness. *Invest Ophthalmol Vis Sci.* 2021 Sep 2; 62(9):2022. doi: 10.1167/19.1021-2020.48. PMID: 34488766. **Günther P, Jank T, Kyma J, Rodriguez FS, Wilker M, Wilker K, Engel C, Strohbl M, Wang M, Franke M, Wilker M, Wehner K, Rauscher FG, Schuster K, Schuster K, Schuster K, Engel C, Thery J, Oberndorf P, Strohbl M, Elze T, Elze T.** Reading position for the eyes: association of retinal nerve fiber layer thickness with cognitive performance in a population-based study. *BMC Commun.* 2023 Nov 8; 3(4):e258. doi: 10.1093/bmccom/mu238. eCollection 2021. PMID: 33520710. **Li D, Rauscher FG, Chu EY, Wang M, Buntmann N, Wilker K, Krause T, Thery J, Engel C, Loeffler M, Elze T, Sotgiu G, Oberndorf P, Strohbl M, Elze T, Wang M, Chu EY, Wang M, Peschel T, Wilker K, Krause T, Thery J, Engel C, Loeffler M, Elze T.** Norms of Intraocular Circumpapillary Retinal Nerve Fiber Layer Thicknesses of 768 Radial Locations. *Transl Vis Sci Technol.* 2020 Aug 12; 9(9):21. doi: 10.1167/tvst.9.9.21. eCollection 2020 Aug. PMID: 32879779. **Krause T, Mariani F, Löffler-Wirth H, Uebel A, Berger C, Strohbl M, Sillig M, Hainel R, Rauscher FG, Schuster K, Peschel T, Hainel H, Wehner J, Zentgraf S, Engel C, Schulz M, Elze T, Löffler M.** The Leipzig Health Atlas - An open platform to research, archive and share biomedical data, analyses and models online. *MedRxiv [preprint].* 2022 Aug 1; doi: 10.1101/2022.08.19.19141985. Epub ahead of print. PMID: 35919777. **Engel C, Wilker K, Zeynalova S, Böhrer B, Binder H, Caporoni U, Erazovsch C, Fuchs M, Hagedorn A, Harzer S, Hinz A, Rauscher FG, Raucha M, Rauscher FG, Schuster K, Sotgiu G, Sotgiu G, Strass A, Thery J, Thery J, Wehner K, Wilker K, Wilker K, Thery J, Schulz M, Loeffler M.** LIFE-Adult-Study working group. Cohort Profile: The LIFE-Adult-Study. *BMJ Epidemiol. 2022* May 28; eyn114. doi: 10.1093/ije/dyab114. Epub ahead of print. PMID: 35642047.

RNFLT(D) - Visualizer

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Browse... No file selected Example_0002_OU_Female_Healthy_SER.pdf pdfs: 1 measurements: 2

id	date	oc	sex	age	diag	bmo	db	valid for
1	0002	2018-04-23	OD	F	66.31	1.3		European Descent (2009) ED-2009, Wang, Elze
2	0002	2018-04-23	OS	F	66.31			European Descent (2009) ED-2009

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RNFL Single Exam Report OU SPECTRALIS® Tracking Laser Tomography

Patient: Example_Female_healthy_FU_LA DOB: 01.Jan.1952 Sex: F
 Patient ID: 0002 Exam.: 23.Apr.2018

Contact: Thomas Peschel: thomas.peschel@imise.uni-leipzig.de
RNFL(D)-Visualizer: <https://apps.health-atlas.de/rnflt-visualizer/>



- Outlook: Improving Models**
- Box-Cox-T, Box-Cox-Green,
 - Box-Cox-Power-Exponential
 - sex dependency
 - ocular laterality dependency
 - higher order dependencies



Retinal layer thickness as a biomarker for monitoring the progression of diseases of the central nervous system –

Medical Data Science using OCT and MRI imaging

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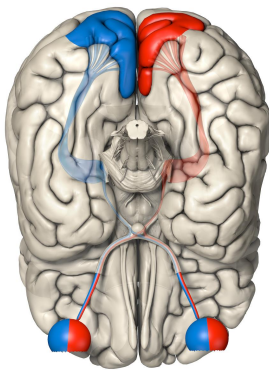
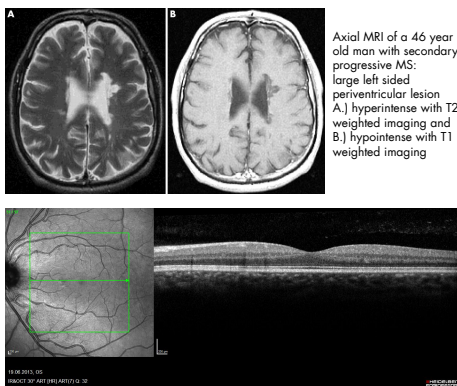
Background

The eye is a window to the brain. There is promising evidence of correlations between retinal layer thickness from optical coherence tomography (OCT) and magnetic resonance imaging (MRI) markers.

→ This results in the potential for science-based clinical application of OCT and MRI in combination for diagnosis and follow-up of various diseases of the central nervous system.

■ In the present study, retinal structures (measured with OCT) are compared with intracerebral structures and pathologies from available MRI examinations. In addition, their correlations to cognitive functions will also be investigated.

■ For this purpose, existing OCT, MRI and data from neuropsychological and cognitive examinations (partly under EEG) of patients of the ZKN and test persons of the ZKN and KJP will be used (retrospective study design).



Methods

complete dataset: 316,681 clinical images from existing documentation of clinical routine for 2,662 individuals (including data and information collected during clinical routine in connection with the OCT and/or MRI examinations)

removal of duplicates

study population: 85,167 clinical images for 2,659 patients of the ZKN as well as for subjects who received OCT and/or MRI examinations at the ZKN and KJP of the University Hospital Carl Gustav Carus Dresden, Germany

exclusion of 4,293 records of 1,047 patients with non-standardised image settings (excluded scans belong to 9 patients who additionally also possess all standardised scans (=records not lost); or to 1,032 patients with only some of the required imaging; and further to 6 patients not included in the standardised sample).

analysis subset: 64,925 clinical images for 1,659 patients imported into the analysis pipeline

- Cross-sectional and longitudinal analyses for three key OCT scan types of the Glaucoma Module by Spectralis spectral-domain Optical Coherence Tomography (SD-OCT), Heidelberg Engineering, Heidelberg, Germany
- Scan settings were:

Macular volume scan with horizontal scan location [PPoleH[APS]] in parallel to the disk-fovea-axis. 61 B-scans with a distance of 120µm between scans inside a 30° by 25° area. High-speed mode with 768 x 496 pixel.

Macular volume scan with vertical scan location [PPoleV[APS]] perpendicular to the disk-fovea-axis. 19 B-scans with a distance of 240µm between scans inside a 30° by 15° area. High-resolution mode with 1536 x 496 pixel.

Circumpapillary ring scan around the optic nerve head [ONHRC[APS]] comprising of three concentric ring scans with 3.5mm, 4.1mm, and 4.7mm diameter (accompanied by 24 line scans). High-speed mode with 768 x 496 pixel.

→ These three scan patterns were matched by regular volume and circular scans of the same resolution settings to increase the number of analyzed patients, when possible.

Results

Standardised SD-OCT imaging at the Multiple Sclerosis Centre Dresden, Centre of Clinical Neuroscience, Department of Neurology, Carl Gustav Carus Faculty of Medicine, Technical University Dresden, Germany was analyzed for fixed scan-settings.

■ Horizontal volume scans with 61 B-Scans, vertical volume scans with 19 B-Scans and one version of circumpapillary Retinal Nerve Fibre Layer scan (either ONH-RC[APS] or one circular B-scan, sometimes combined with 24 radial scans) were present in combination in 1,659 patients for 1,640 right eyes and 1,600 left eyes.

■ Of those, 1,379 (right eye) and 1,347 (left eye) presented with the ideal imaging by the Glaucoma Module (PPoleH[APS], PPoleV[APS] and ONH-RC[APS]). For 422 right eyes and 408 left eyes comparable scan combinations were obtained.

■ In total, 64,925 records of a final sample of 1,659 patients were imported into the analysis pipeline. This clinical dataset consists of routine visits of 1,630 patients (60,233 records) as well as 53 specific study patients with 4,702 records (7% of records).

Image Type	B-Scans
Set 1 OCT Volume Scan	61
Set 2 OCT Circle Scan	1
Set 3 OCT Volume Scan	19
Set 4 OCT Radial Scan	24
Set 5 OCT Radial Circle Scan	27

Exam Date freq	Legend:
Group 1 # <= 2	R1: number of eyes with full set of records for Set 1, Set 3 and Set 5;
Group 2 2 < # <= 4	L1: number of eyes with full set of records for Set 1, Set 3 and Set 5
Group 3 4 < # <= 6	
Group 4 6 < # <= 8	
Group 5 8 < # <= 10	
Group 6 10 < #	

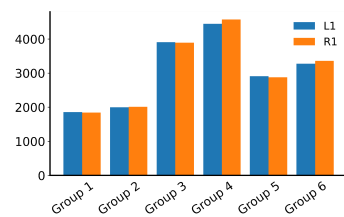
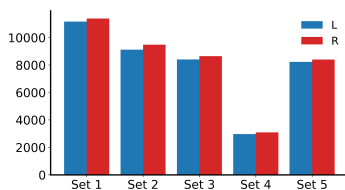


Figure 1 Frequency of desired image settings; Legend: R: right eye; L: left eye

Figure 2 Frequency of examinations with the combination of desired settings: per eye

Discussion

At ZKN, in combination with joint studies by ZKN and KJP, 80,874 images were obtained in the aforementioned volume and circular scan settings, Set 1, Set 3, and Set 5 or comparable (right eye: 40,985; left eye: 39,889). This points to a very high level of standardisation of imaging in this clinical setting. This unprecedented dataset is of high value to the research questions of ZKN and KJP and will serve as the ideal basis for cross-sectional and longitudinal analyses of diseases of the central nervous systems. A substantial number of patients had standardized imaging by the Spectralis Glaucoma Module, for the desired scan settings, 64,925 records of 1,659 patients were available. Inside this sample, 53 specific study IDs were counted, some of which could be the topic of a separate analysis.

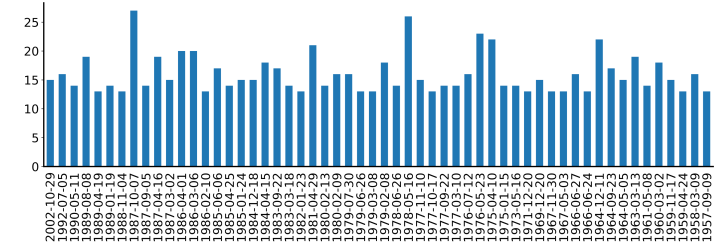


Figure 3 Frequency of exams per patient in Group 6, sorted by birthdate for longitudinal analysis;

Outlook: The 1,659 patients with available 64,925 OCT records will be stratified into clinical research questions to focus on cross-sectional and longitudinal analyses of diseases of the central nervous system, this will include data from MRI and other clinical information. The research will centre on Multiple Sclerosis (MS), with investigations on disease progression, and therapy, with a specific focus on relapsing and remitting MS.

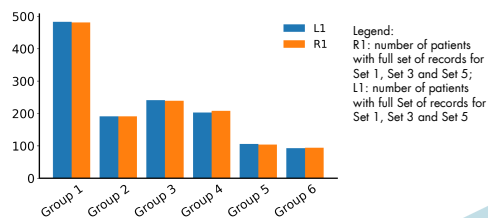
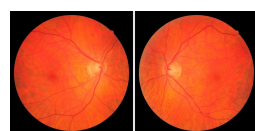


Figure 4 Number of patients with frequency of examinations with the combination of desired settings





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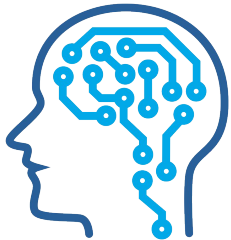


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
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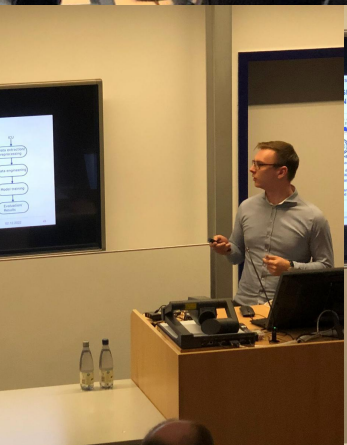
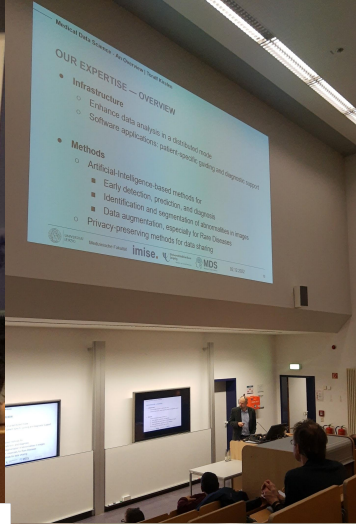
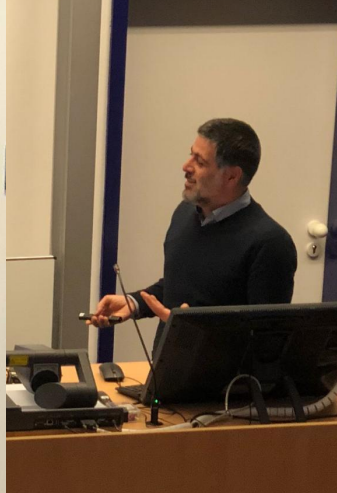
Leipzig, 02.12.2022



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Medical Data Science

Veranstaltung
**"Medical Data Science – Linking
 Medicine and Data Sciences"**
 Prof. Dr. Toralf Kirsten
 02.12.2022 14.30 - 15.30 Uhr Hörsaal,
 Liebigstraße 12




RESEARCH PROFILE

- Semantic integration of medical data
- Privacy-preserving distributed analysis of medical data
- Applications of novel artificial intelligence methods in medicine and healthcare
- Generation of synthetic data for medicine and healthcare applications
- Current application domains:
 - Leukodystrophies (rare diseases)
 - Ophthalmology
 - Diabetes

We are open to new projects, topics, and ideas in the field of medical data science. Please do not hesitate to contact us for more information if you are interested in collaborating with us or have any questions.



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WE INTRODUCE OURSELVES

We are an interdisciplinary team of Medical Data Science at Leipzig University Medical Informatics Center. We conduct innovative research at the intersection of medical informatics and patient care using artificial intelligence and medical data science to increase benefits for patients, the healthcare system, and medical research. Our focus is twofold: First, we develop a secure computing infrastructure for the preparation, processing and analysis of medical data. This improves the accessibility of such data for research purposes, as medical data are often available from multiple locations, for example in distributed computing. Second, we perform medical data analysis and modeling to advance medical research and also support clinicians. For this, we employ both classical statistical methods and state-of-the-art machine and deep learning techniques.



FAIR DATA

Knowledge derived from captured and collected data has become increasingly important in research and development. In order to reduce difficulties in data sharing in this respect and maximize the value from data, data and metadata should comply with the FAIR principles - Findable, Accessible, Interoperable, and Reusable. To this end, we develop data structures and standardize the process for accessing the data to ensure that the basic requirements for reproducibility are met. We are part of the National Research Data Infrastructure (NFDI4Health), which is an initiative in this context.



INFRASTRUCTURES AND ALGORITHMS FOR DISTRIBUTED ANALYTICS

Personal medical data are often collected and available in multiple locations. Due to privacy concerns and the potentially large volume of medical records, the pooling concept for centralized analysis may not work. As part of the international Personal Health Train (PHT) network, we are working with our collaborators to develop an infrastructure for secure distributed computing. Such PHT infrastructure adheres to the principle of “bring-the-analysis-to-the-data”. We have successfully implemented this infrastructure in several research projects within the medical field.



AI-BASED MODELING OF LEUKODYSTROPHIES

Leukodystrophies are a family of rare diseases with an estimated incidence of less than 1:40,000 worldwide. Patients with leukodystrophy typically experience an Odyssey and undergo a challenging, expensive and time-consuming journey. An important issue is that the disease is commonly misdiagnosed and therefore incorrectly treated. Furthermore, since only a few cases are known, there is a lack of validated treatment guidelines. In close collaboration with the Myelin Center at the Leipzig University Medical Center and other interdisciplinary international consortia, we frequently make use of a variety of artificial intelligence methods and analyze data from leukodystrophy patients to support earlier diagnosis, for example.



ABOUT TORALF KIRSTEN

Prof. Toralf Kirsten heads the Medical Data Science department. With his broad computer science background, which ranges from the conception and implementation of IT infrastructures to methods and algorithms of data analysis, he is involved with the department in various national and international research projects. He is actively involved in the Medical Informatics Initiative (MII), the Network University Medicine (NUM), the National Research Data Infrastructure (NFDI) and the international GOFAIR Initiative, which interacts closely with the European Open Science Cloud (EOSC).

