

# DeepSpA

Deep Speech Analysis for Clinical Trials

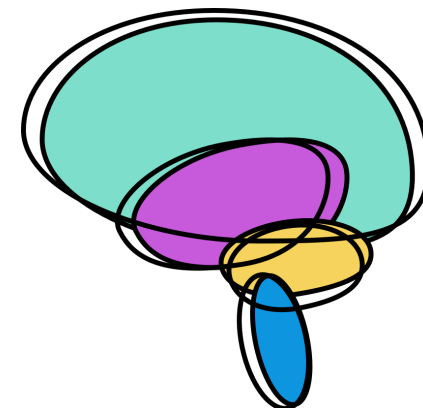
—Novel Digitalized Markers for Screening & Monitoring of Patients—



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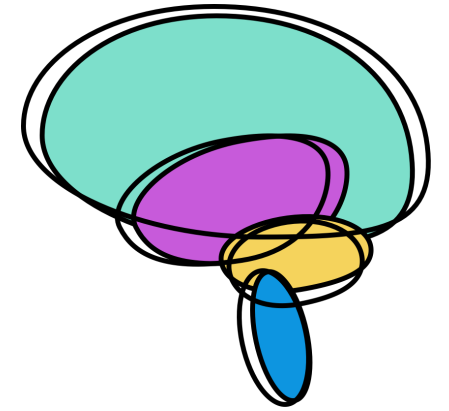
# A X-KIC EIT-Health & -Digital project:



- European Institute of Innovation & Technology (EIT)
  - Fully funded
  - Builds on a previous EIT Digital project and existing assets from partners
- January 2019 until June 2020 (18 months)
- 6 partners:
  - **Janssen**
  - **inria** (Institut national de recherche en informatique et en automatique)
  - **DFKI** (German Research Centre for Artificial Intelligence)
  - **Maastricht UMC+** (Maastricht University Medical Center+)
  - **AIA** (Association Innovation Alzheimer)
  - **ki elements** (German AI start-up)

# Deep Speech Analysis for Clinical Trials in AD—Why care?

# Think about a new era of clinical trials: e.g. secondary prevention of AD

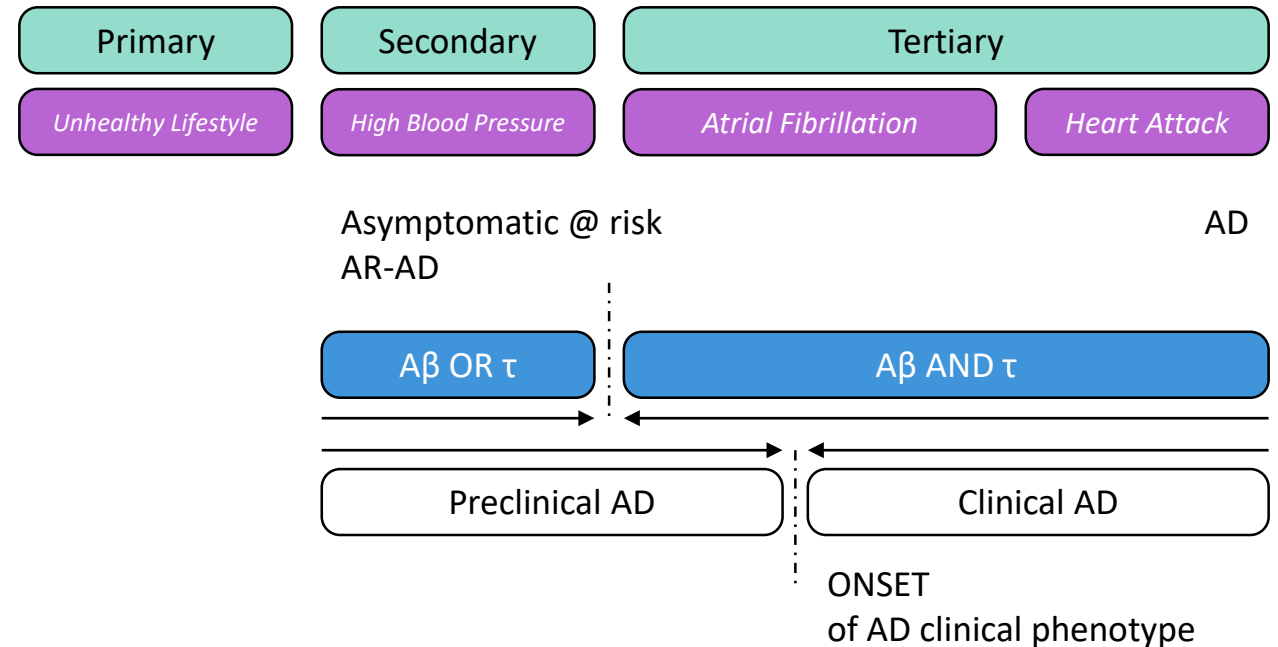


- The A4 Study

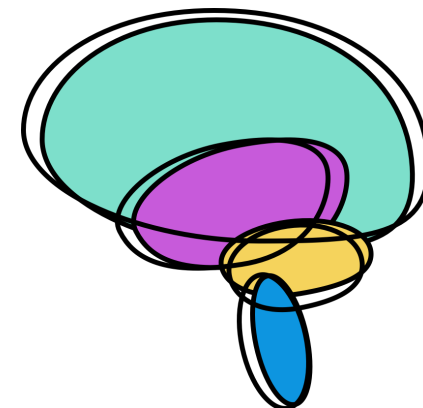
(Sperling et al., 2014) → led by Lilly

- The European Prevention of Alzheimer's Dementia (EPAD) Project

(Ritchie et al., 2016) → coordinated by Janssen



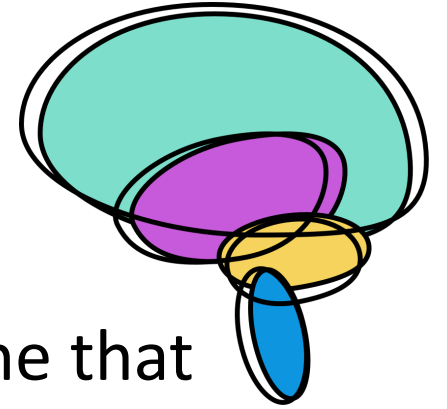
# The challenge: onboarding the relevant cohort



- Phenotype:
  - A $\beta$ +
  - no clinical symptoms in neurocognition as compared to traditional cross-sectional norms
    - MMSE ~30
    - CDRS ~0
- Trial inclusion in e.g. EPAD:
  - Include participants through large EU cohorts (often epidemiological or cardio-vascular cohorts)
  - Inclusion rate 1/100 –1/500
  - f2f assessments in clinical facilities → ~8k\$ per participant screening costs

DeepSpA's goal is to develop an automated, remote pre-screening and monitoring solution for clinical trial use cases focusing on cognitive decline.

# Reaching the overall goal:

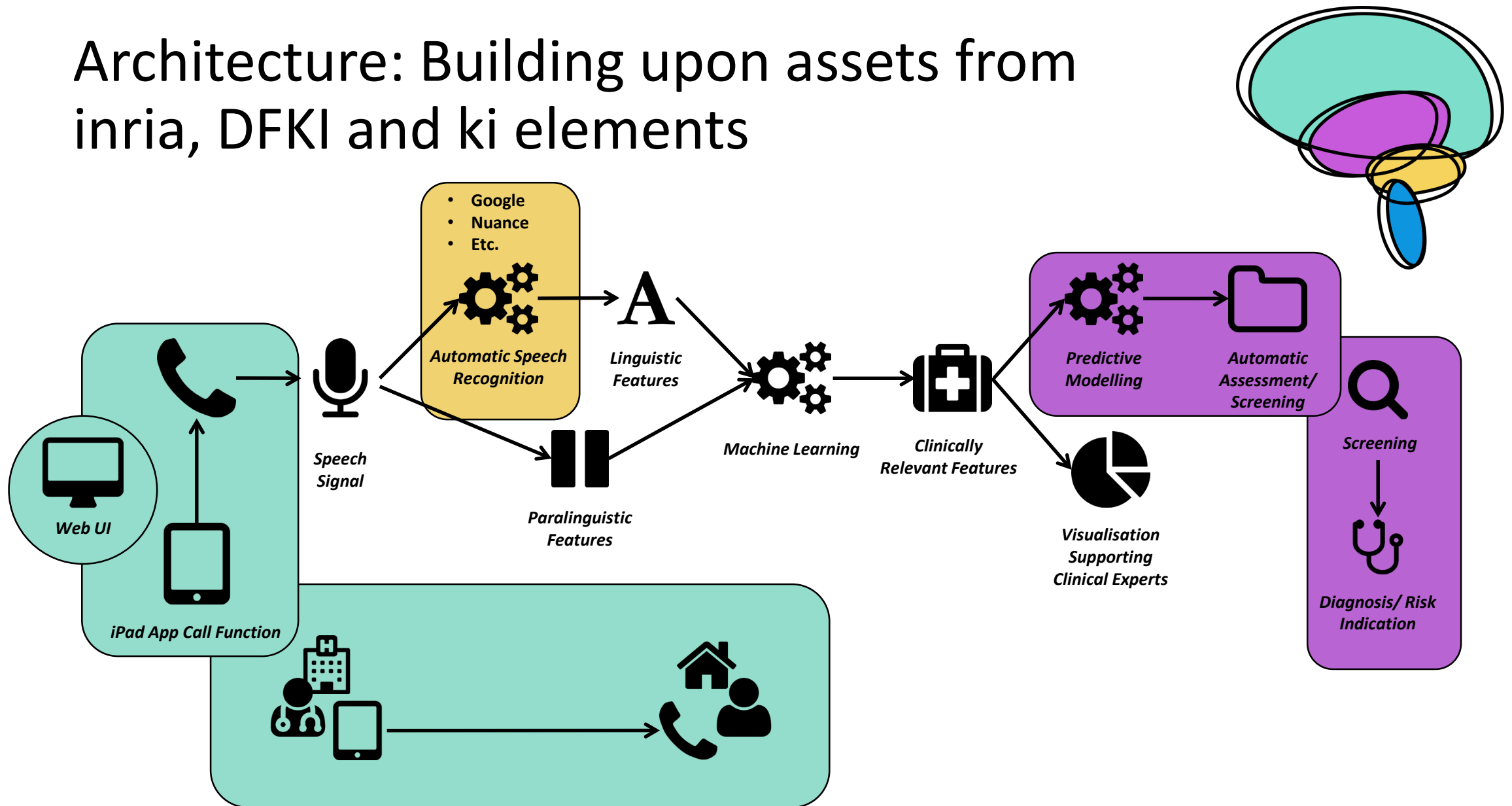


1. Define predictive digital biomarkers for cognitive decline that only need an ordinary telephone for collection
2. Implement a solution that allows for remote (telecommunication-based) assessment and risk classification
3. Prove the technical feasibility of the remote scenario as compared to the traditional face-to-face one
4. Prove the scientific & economic validity of the classification algorithm: e.g. detection of at-risk cognition above today's inclusion rates

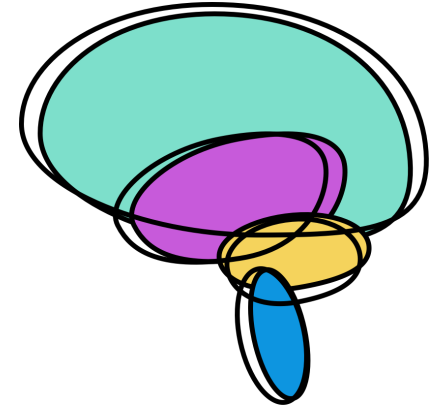
Technical challenge: Is phone-based longitudinal cognitive pre-screening and monitoring actually feasible?



# Architecture: Building upon assets from inria, DFKI and ki elements



# Feasibility study for the remote use case

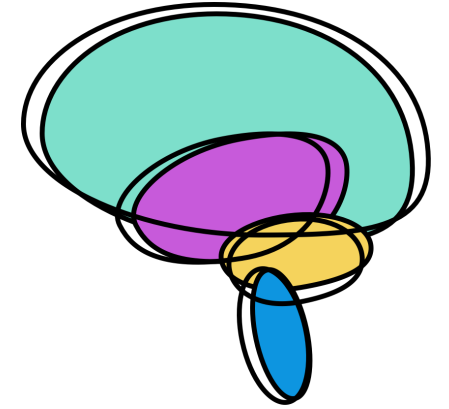


Phone assessment for pre-screening

- Enabling remote phone-based pre-screening: can we use speech only cognitive tests for automatic AR-AD detection?
- Maastricht (NL) feasibility study based on BioBank Alzheimer Center Limburg Cohort
- Phone-based tests

# The Maastricht study

*60 participants, Age ~ 60, Subjective Cognitive Decline, Biobank  
Alzheimer Center Limburg Cohort*



## Baseline F2F assessment 1

- Audio Recordings Cognitive Tests
  - Verbal Learning Test (V1)
  - Verbal Fluency (V1)

+ biomarkers (gold standard)  
+ consents

## Semi-automated telephone call

+ verbal learning (V2)  
+ digit span  
+ verbal fluency  
+ questionnaire SCI

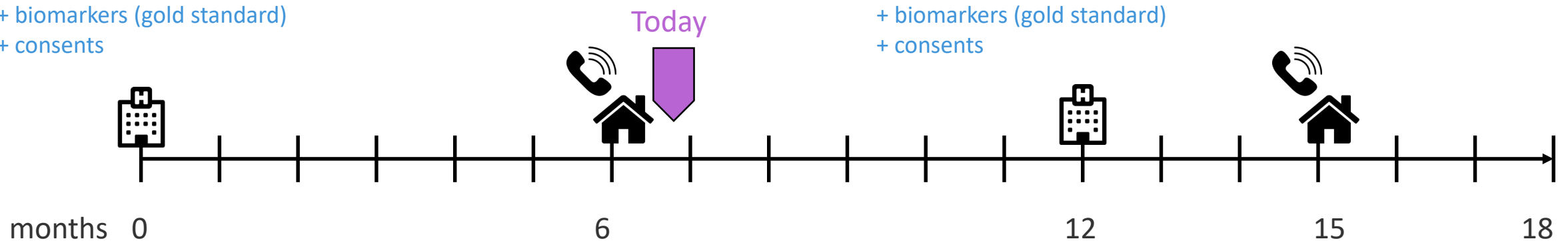
## Baseline F2F assessment 2

- Audio Recordings Cognitive Tests
  - Verbal Learning Test (V3)
  - Verbal Fluency (V2)

+ biomarkers (gold standard)  
+ consents

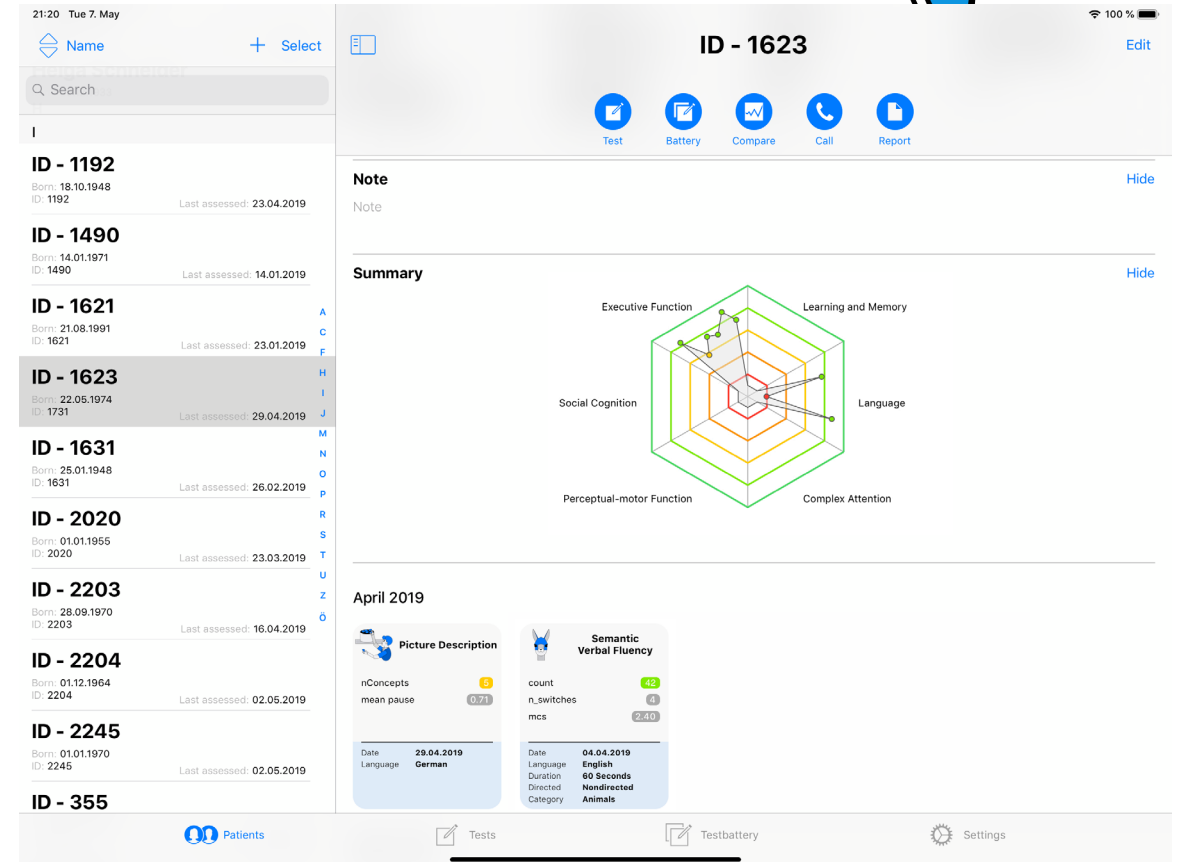
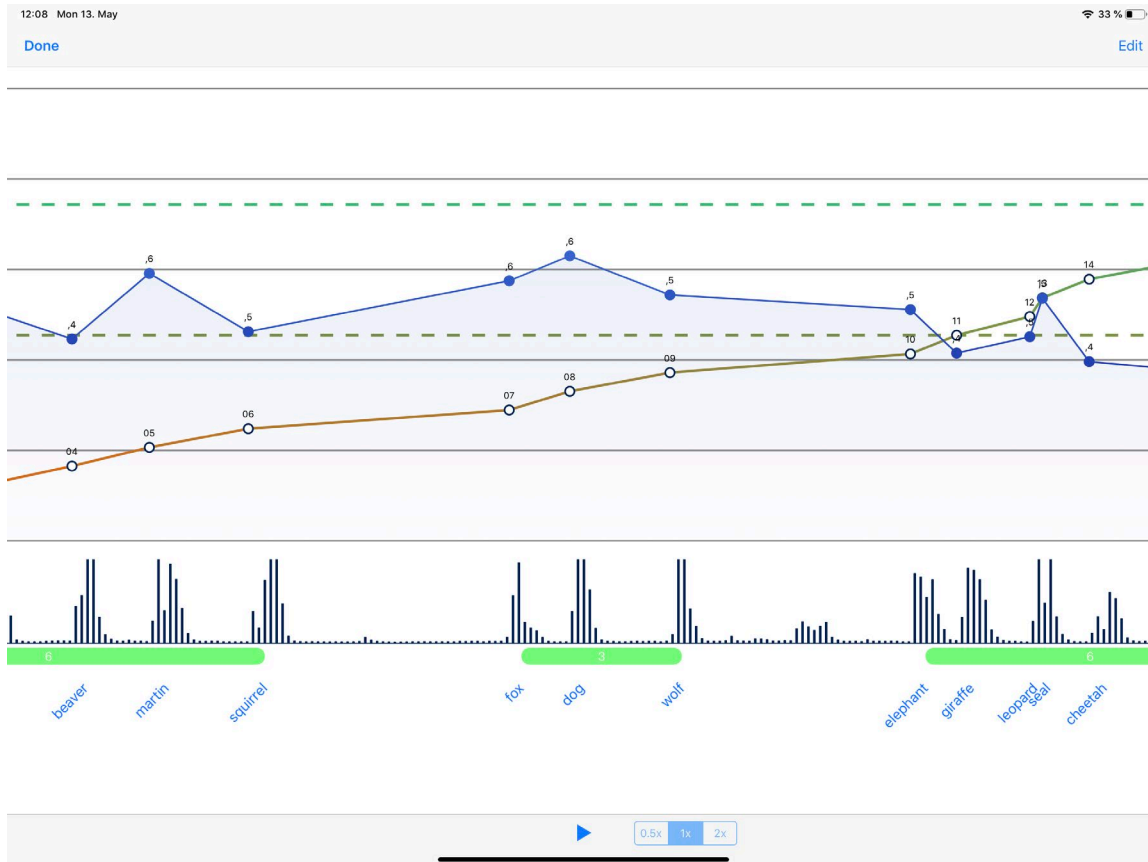
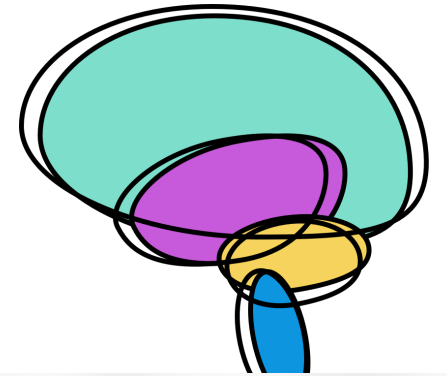
## Fully-automated telephone call

+ verbal learning  
+ verbal fluency



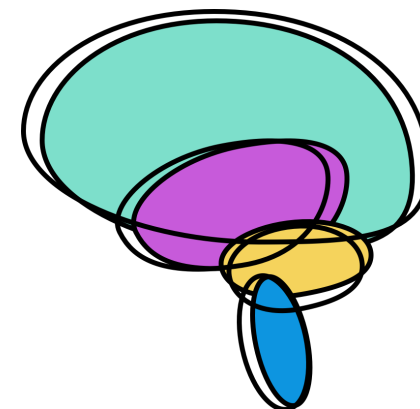
# Building upon the $\Delta$ elta platform

*An iPad app that digitalises and enhances classical neuropsychological assessment using Artificial Intelligence (AI). With  $\Delta$ elta, assessment becomes more standardized, smarter and faster.*



Scientific challenge: Can we  
build an algorithm that  
effectively screens for the  
relevant population?

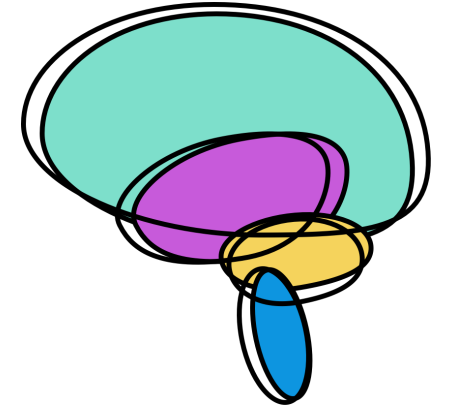
# Can we train a speech-based classification algorithm?



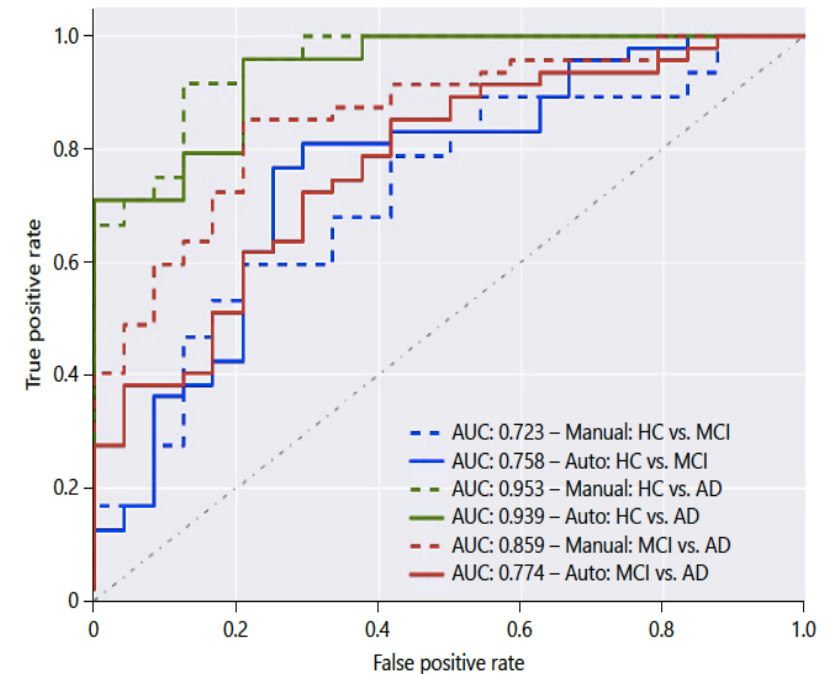
Clinical Symptoms (including cognition)	
	+
	-
Biomarker (e.g. A $\beta$ )	+
	-
	X

- Detect clinical symptoms (the soft problem)
  - Detection of clinical cognition is our classic research case (König et al., 2018; Tröger et al., 2018; Linz et al., 2017a & b;...)
- Biomarker-related early cognitive changes (the hard problem)
  - The challenge: for training screening models, we need the relevant population
  - Looking for opportunities: training on longitudinal well-phenotyped cohort

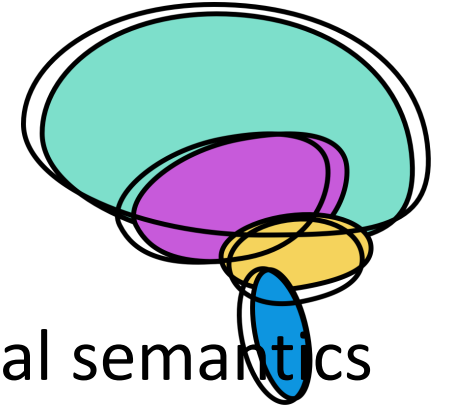
# Scientific evidence I



- Fully automated screening based on speech analysis
  - An automated pipeline analyses the semantic verbal fluency (SVF) with the same result as a classification on manually transcribed data by humans ([König et al., 2018](#))
  - A Phone based automatic SVF analysis is valid compared to human raters ([Tröger et al., 2018](#))
  - Dementia screening and staging scores (MMSE, CDR-SB) can be predicted from advanced SVF features using machine learning ([Linz et al., 2017a](#))



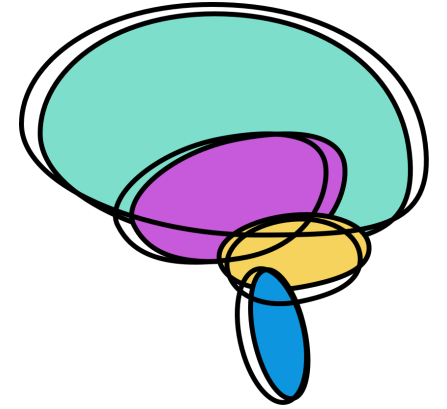
# Scientific evidence II



- Model of neurocognitive functions through computational semantics
  - automatic semantic clusters and switches as measures for semantic memory retrieval and executive control processes ([Tröger et al., 2019](#); [Linz et al., 2017b](#))
  - computational modelling of the SVF serves as predictor ([Linz et al., 2018](#))
- Analysis of additional para-linguistic (e.g. pauses, pitch, etc.) and content features
  - Affective syndromes related to AD can be characterized automatically using para-linguistic speech analysis ([König, et al., 2019](#)) and sentiment psycholinguistic analysis ([Linz et al. 2019](#))

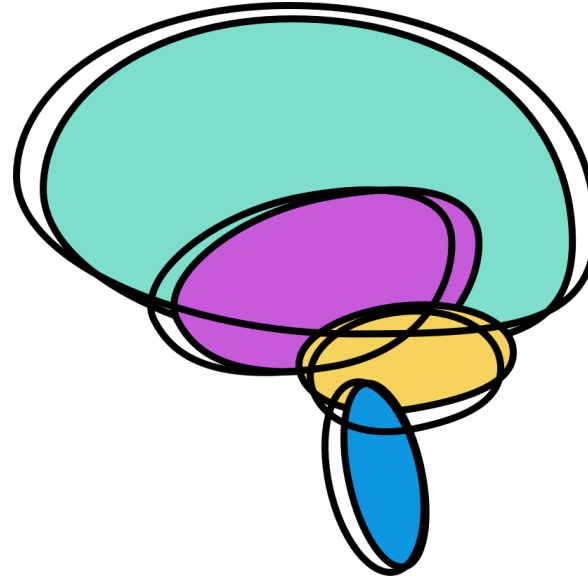


# Unique Differentiators



We work on a solution that:

1. scales easily to multiple languages,
2. is cost-efficient because of a (semi-) automated workflow,
3. has no technical barrier (ordinary telephone is enough) and
4. uses existing clinical tests/procedures; a clinician can always be in the loop and knows perfectly well how to interpret results!



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