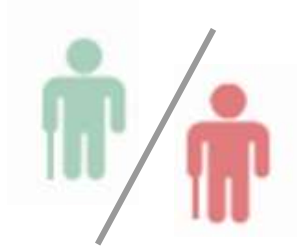


ML in healthcare

Christian Salvatore
Scuola Universitaria Superiore IUSS Pavia

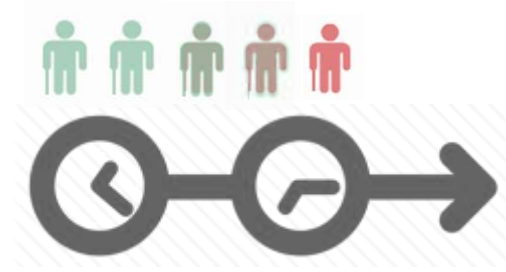
christian.salvatore@iusspavia.it

Machine learning applied to medical imaging



Diagnosis (early/differential)

is the patient healthy?



Prognosis

what will be the course of the disease?



Screening

*are the patients -within
a population- healthy?*



Treatment addressing

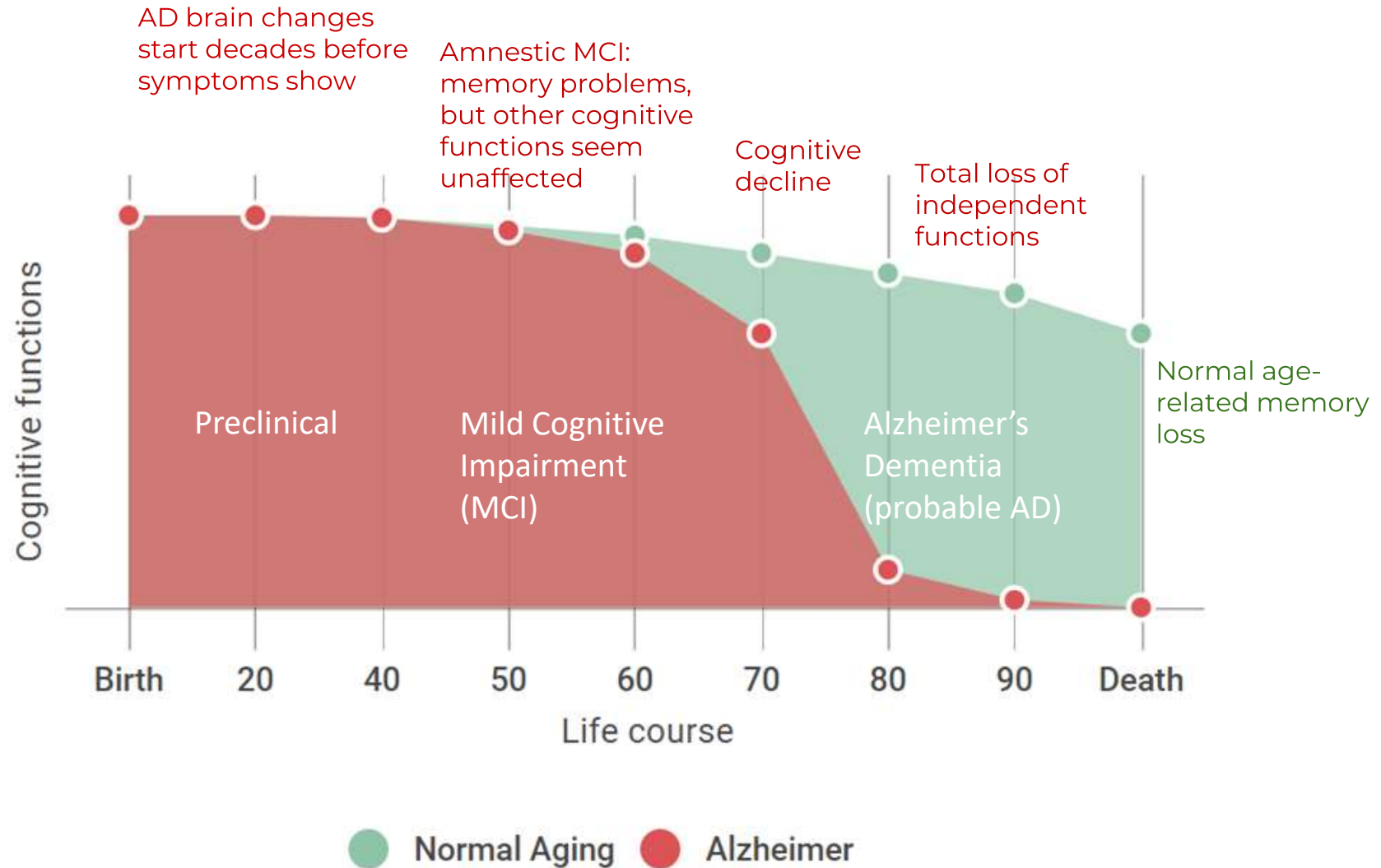
*will this therapy be effective
for this patient?*

ML Models | Examples

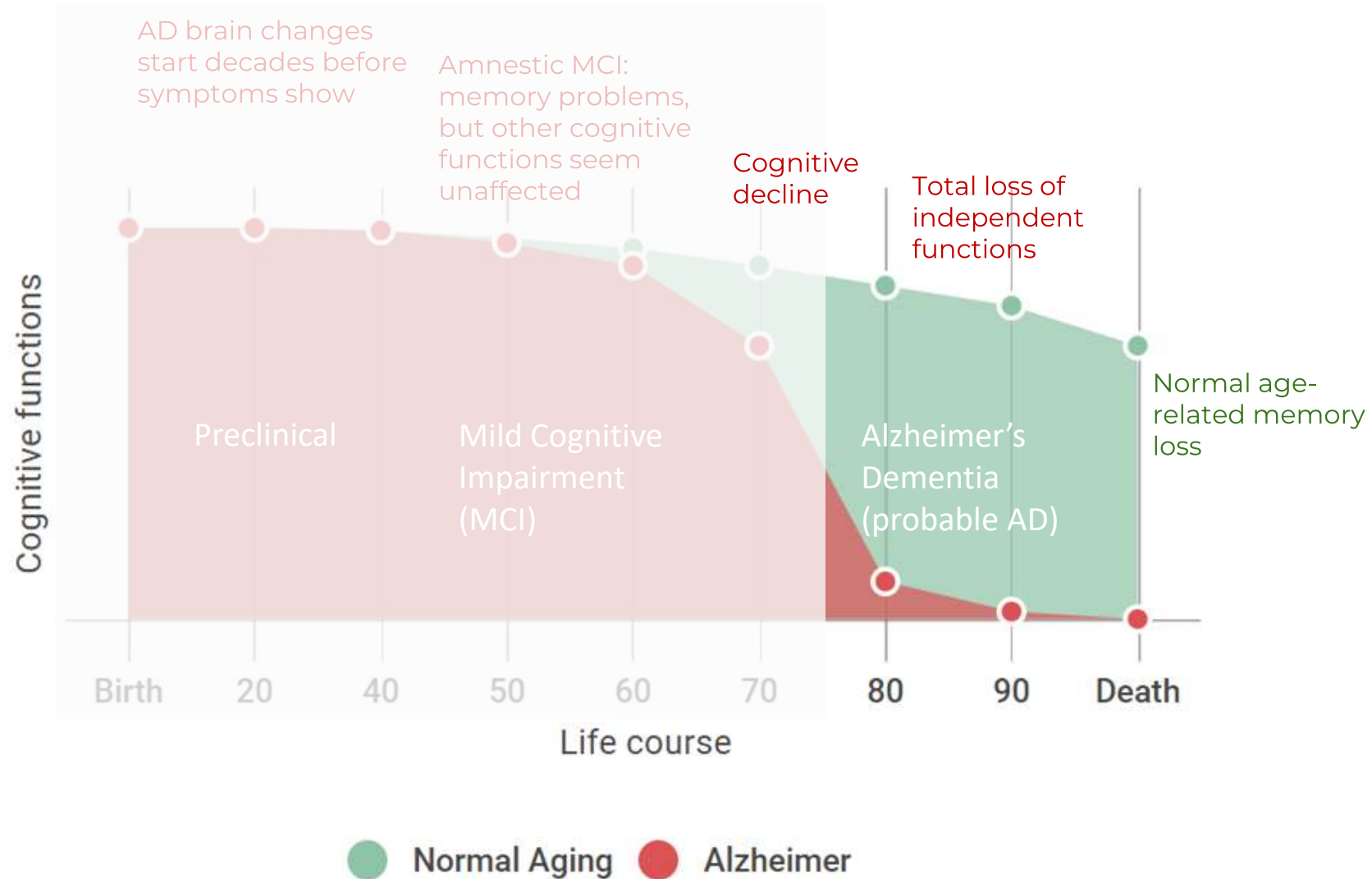
(using different types of data)

Alzheimer's Disease

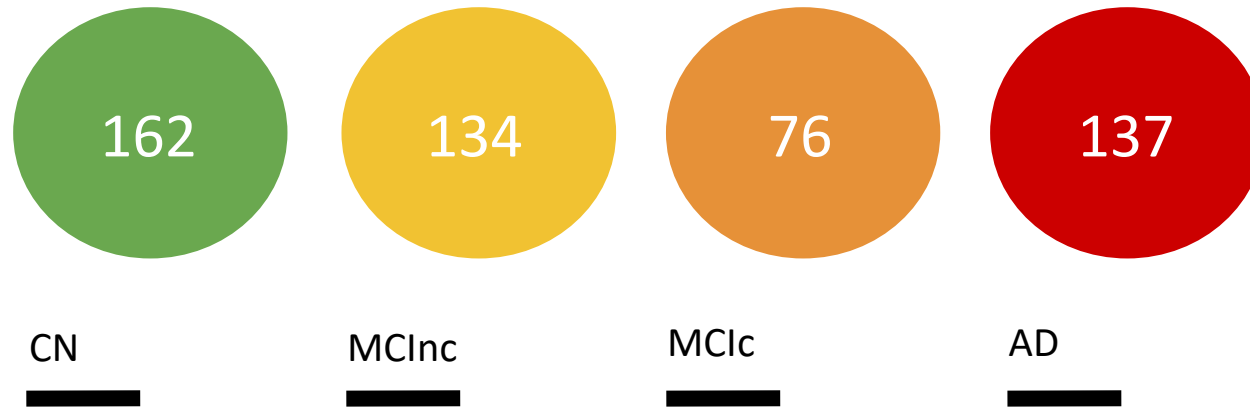
ML models | Examples



ML models | Examples



Clinical Diagnosis of Alzheimer's Disease



AD
MCIC
MCInc
CN

Alzheimer's Disease

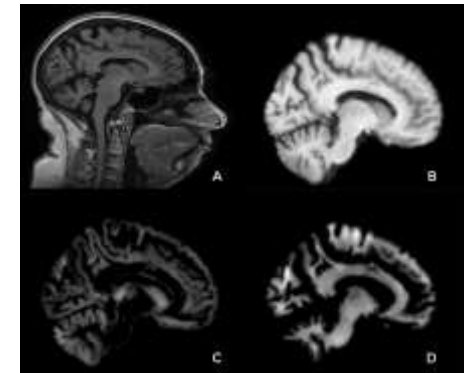
Mild Cognitive Impairment, converting to Alzheimer's Dementia

Mild Cognitive Impairment, not converting to Alzheimer's Dementia




Cognitively-Normal subjects

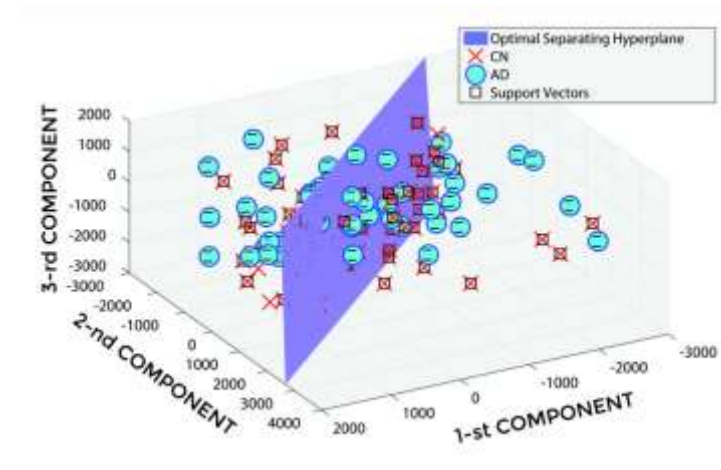
A dataset
of 509 subjects

Structural MRI
T1 weighted
1.5 Tesla

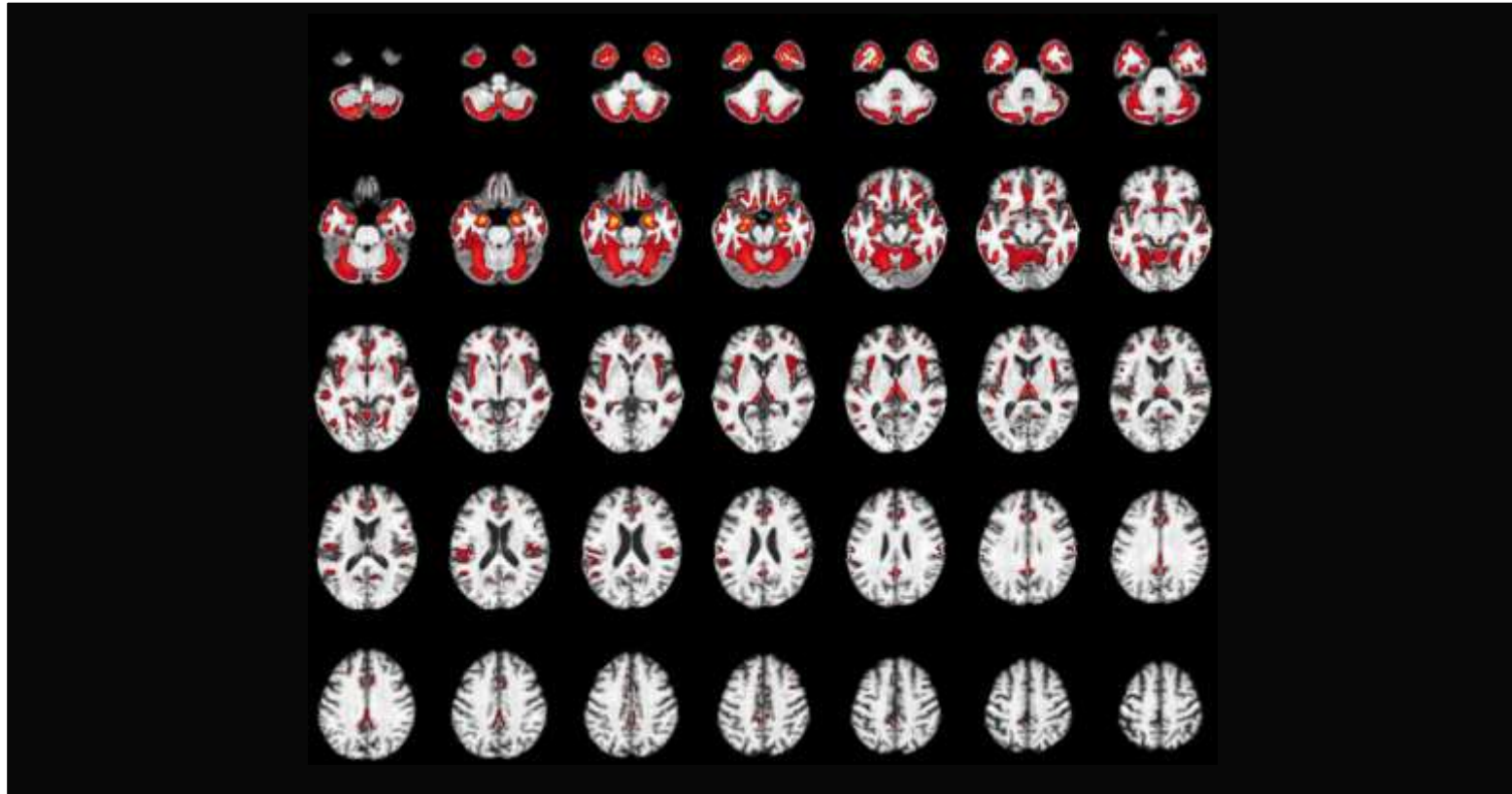


Magnetic resonance imaging biomarkers for the early diagnosis of Alzheimer's disease: A machine learning approach.
Salvatore et al. 2015, Frontiers in Neuroscience.

	Clinical Diagnosis			Accuracy (automatic classification)
	AD	vs	CN	76%
	MCIc	vs	CN	72%
	MCIc	vs	MCIc	66%



Magnetic resonance imaging biomarkers for the early diagnosis of Alzheimer's disease: A machine learning approach.
Salvatore et al. 2015, Frontiers in Neuroscience.

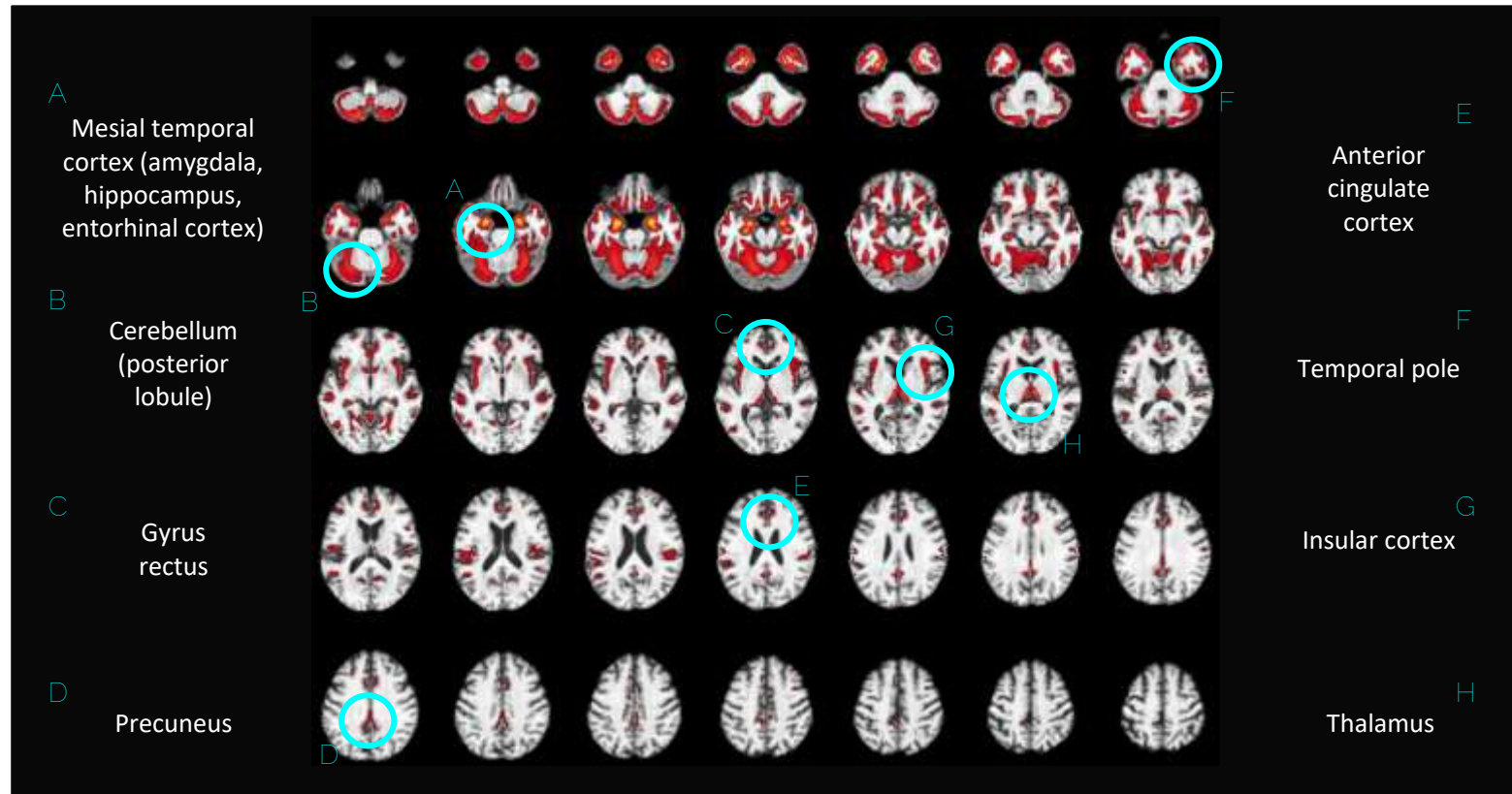


**Best
Structural-MRI
Predictors**

AD vs CN

Magnetic resonance imaging biomarkers for the early diagnosis of Alzheimer's disease: A machine learning approach.
Salvatore et al. 2015, Frontiers in Neuroscience.

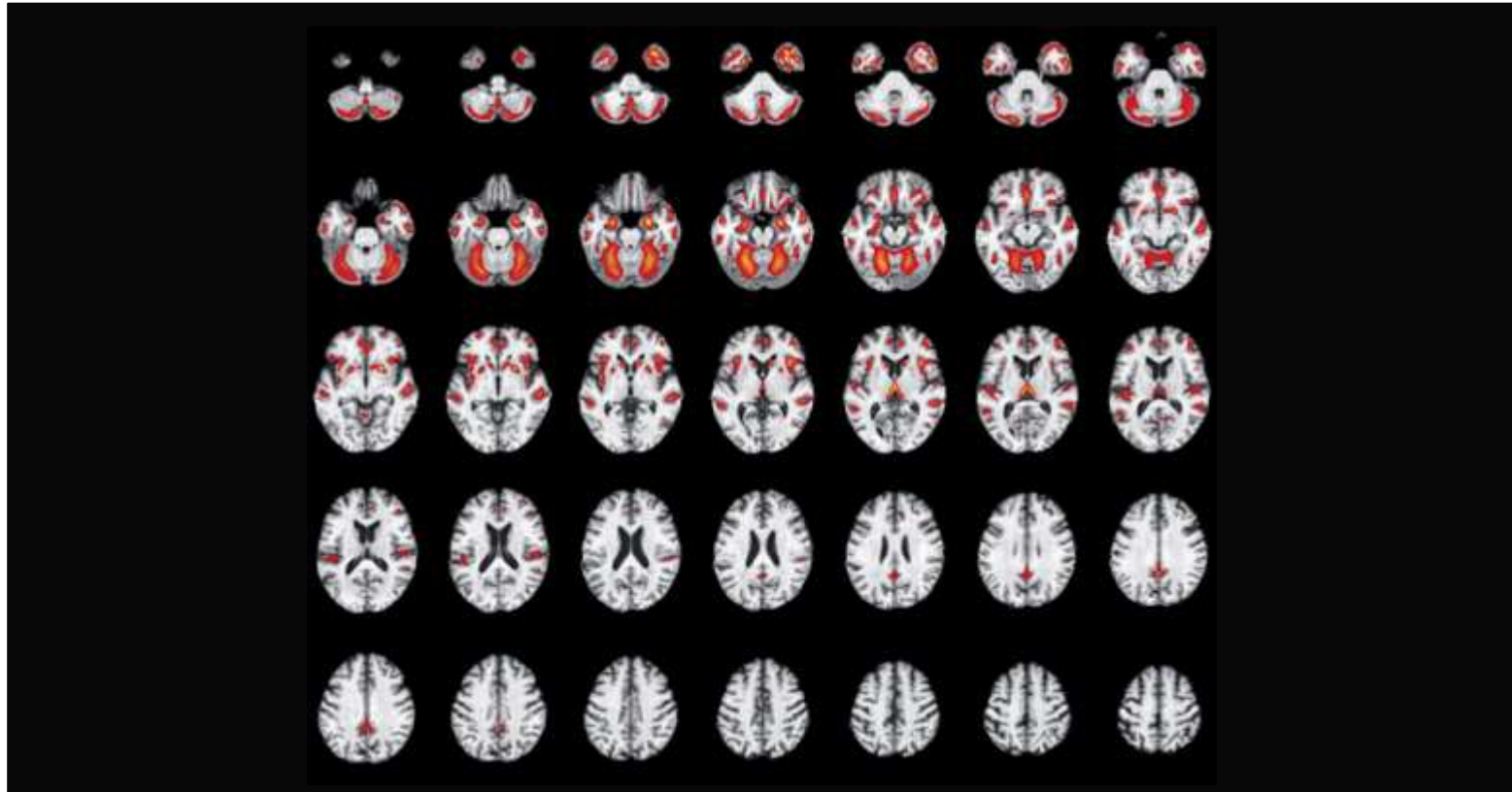
ML models | Examples



**Best
Structural-MRI
Predictors**

AD vs CN

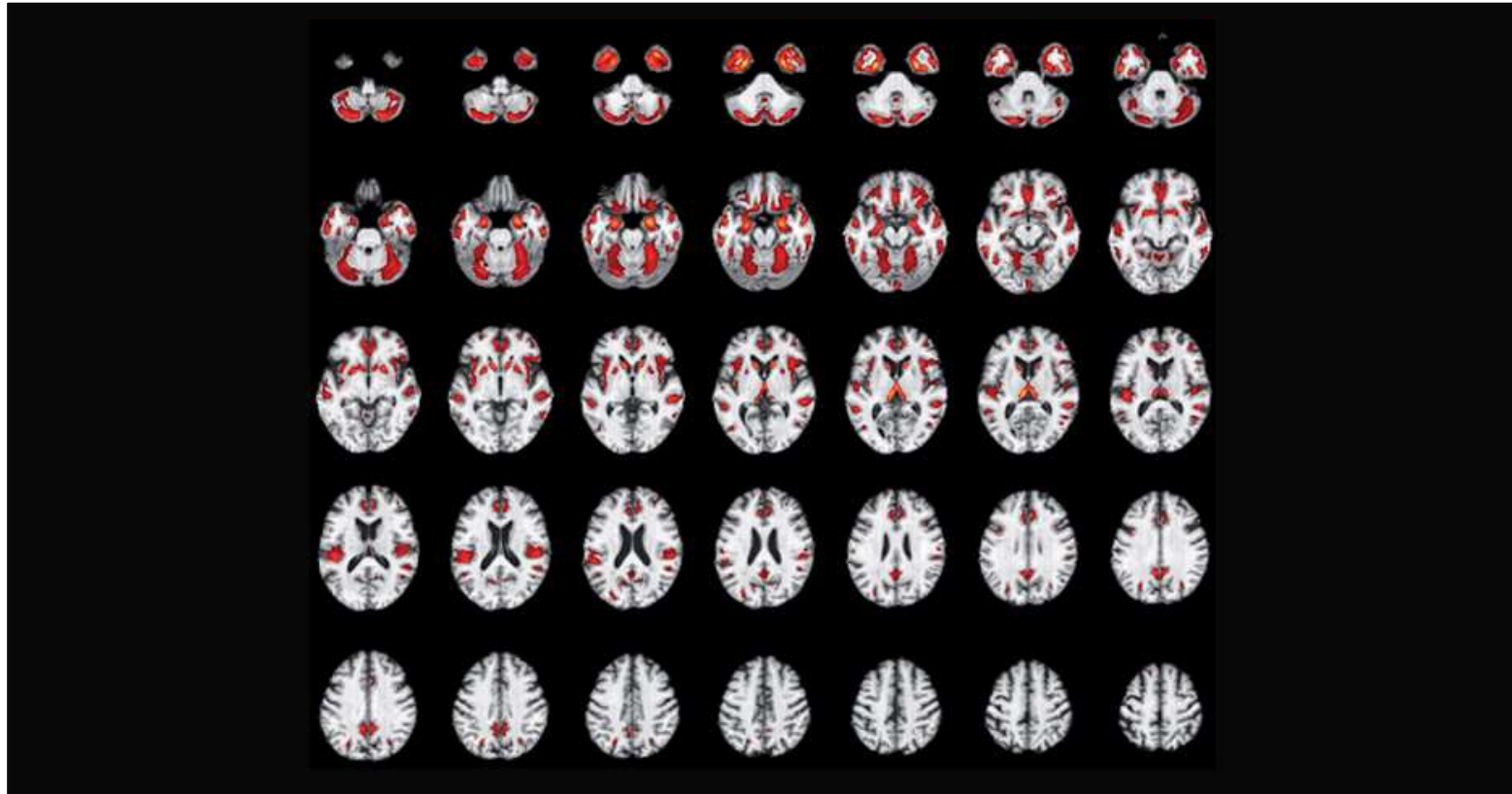
Magnetic resonance imaging biomarkers for the early diagnosis of Alzheimer's disease: A machine learning approach.
Salvatore et al. 2015, Frontiers in Neuroscience.



**Best
Structural-MRI
Predictors**

MCIc vs CN

Magnetic resonance imaging biomarkers for the early diagnosis of Alzheimer's disease: A machine learning approach.
Salvatore et al. 2015, Frontiers in Neuroscience.

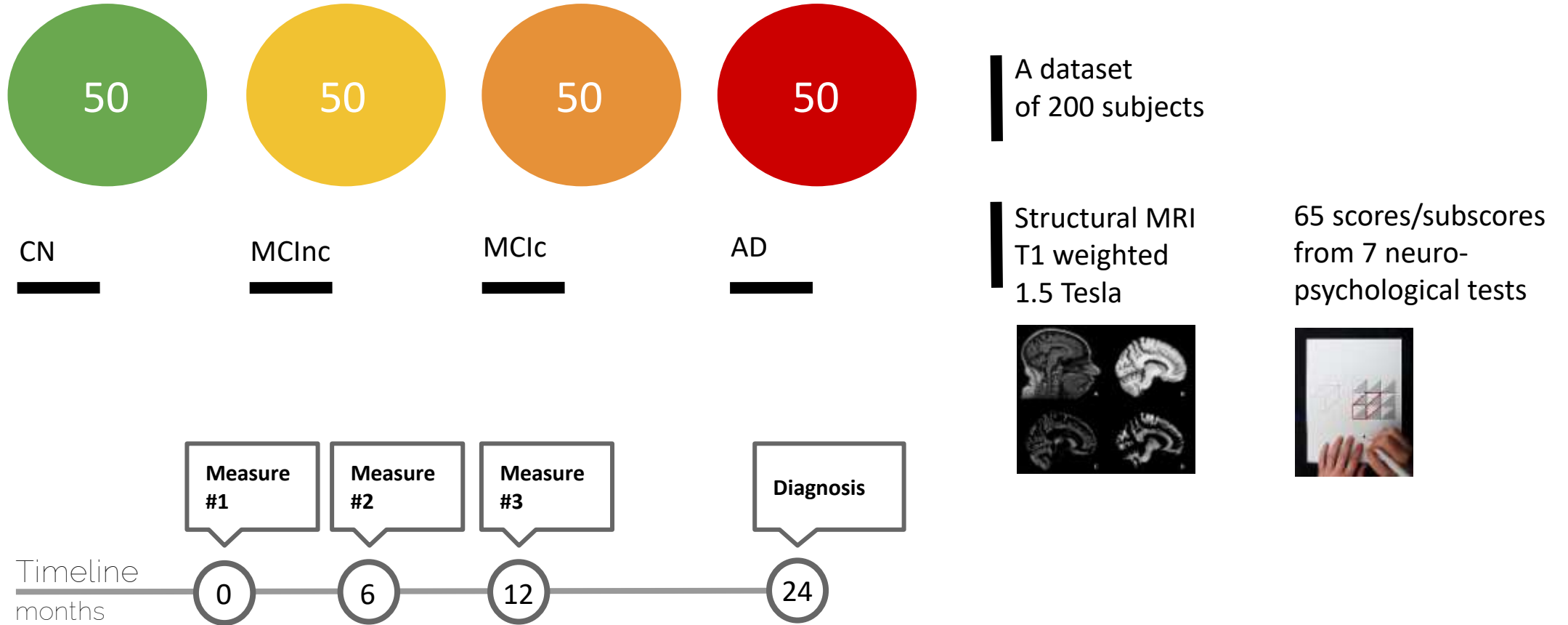


**Best
Structural-MRI
Predictors**

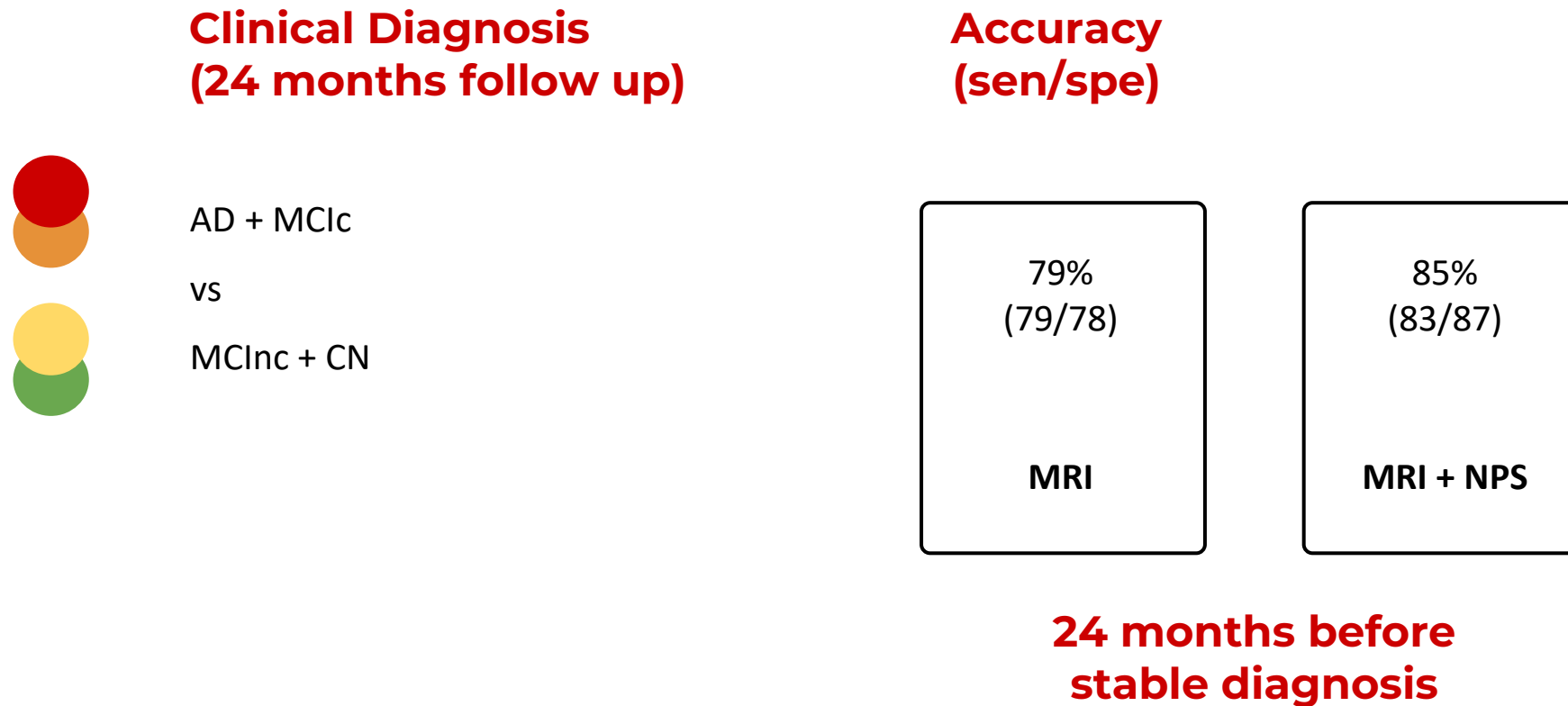
MCIc vs MCIc

Magnetic resonance imaging biomarkers for the early diagnosis of Alzheimer's disease: A machine learning approach.
Salvatore et al. 2015, Frontiers in Neuroscience.

Clinical Diagnosis of AD at 24 months follow up



MRI characterizes the progressive course of AD and predicts conversion to Alzheimer's dementia twenty-four months before probable diagnosis. Salvatore et al. 2018, *Frontiers in Aging Neuroscience*.



MRI characterizes the progressive course of AD and predicts conversion to Alzheimer's dementia twenty-four months before probable diagnosis. *Salvatore et al. 2018, Frontiers in Aging Neuroscience.*

ML models | Examples

Best Neuropsychological Predictors

24 months
before
stable
diagnosis

Ability in remembering appointments, family occasions, holidays, medications in [FAQ](#)
Functional abilities

Ability in writing checks, paying bills, or balancing checkbook in [FAQ](#)
Functional abilities

Ability in assembling tax records, business affairs in [FAQ](#)
Functional abilities

Total score of trial 5 in [AVLT](#)
Memory and learning

Ability in keeping track of current events in [FAQ](#)
Functional abilities

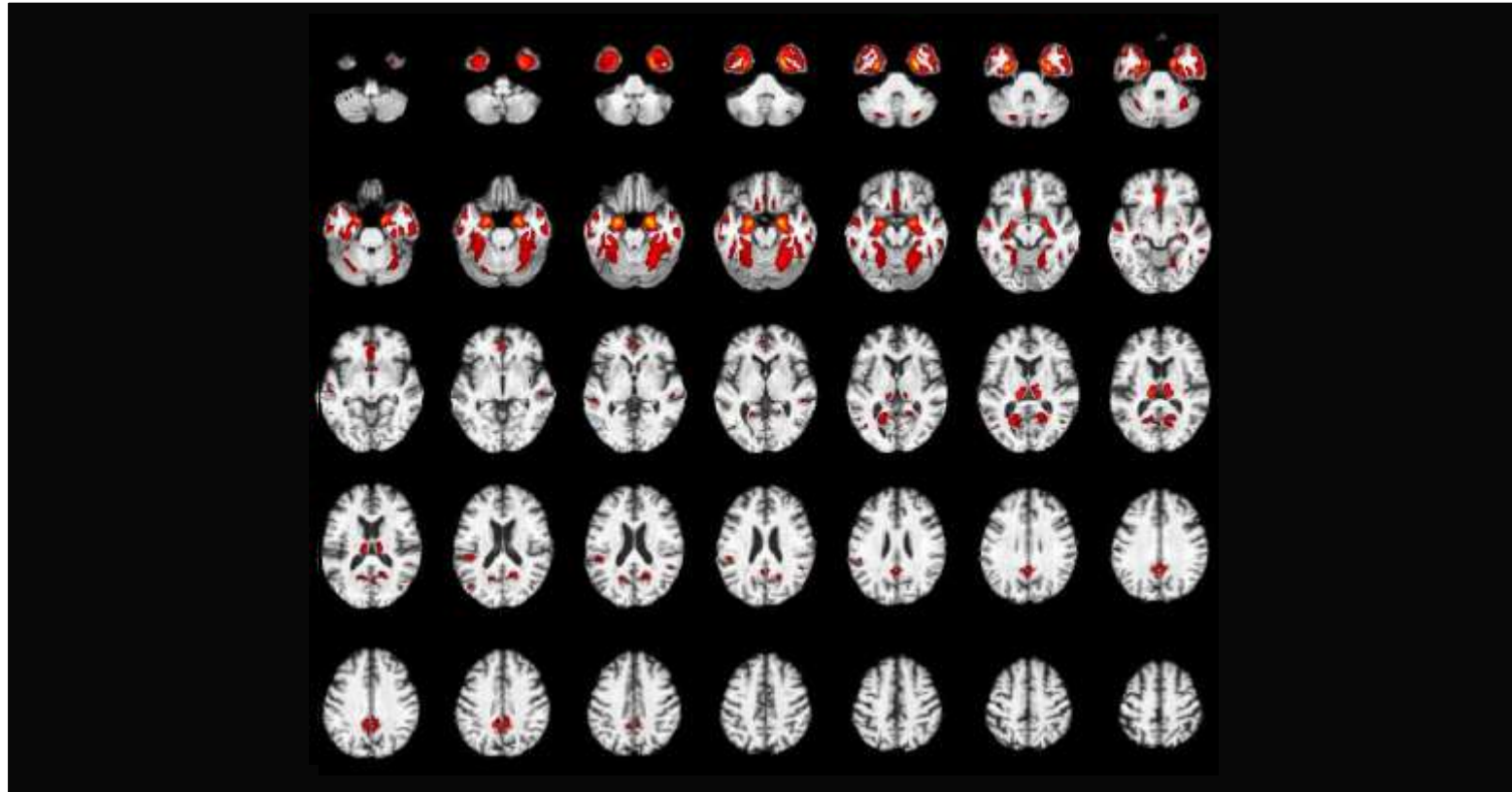
Total intrusions of trial 1 in [AVLT](#)
Memory and learning

Correct answers in the Backwards task in [Digit-Span Test](#)
Working memory

Correct answers in Vegetables task in [Category Fluency Test](#)
Language

Correct answers after a 30-min delay in [AVLT](#)
Memory and learning

MRI characterizes the progressive course of AD and predicts conversion to Alzheimer's dementia twenty-four months before probable diagnosis. Salvatore et al. 2018, *Frontiers in Aging Neuroscience*.

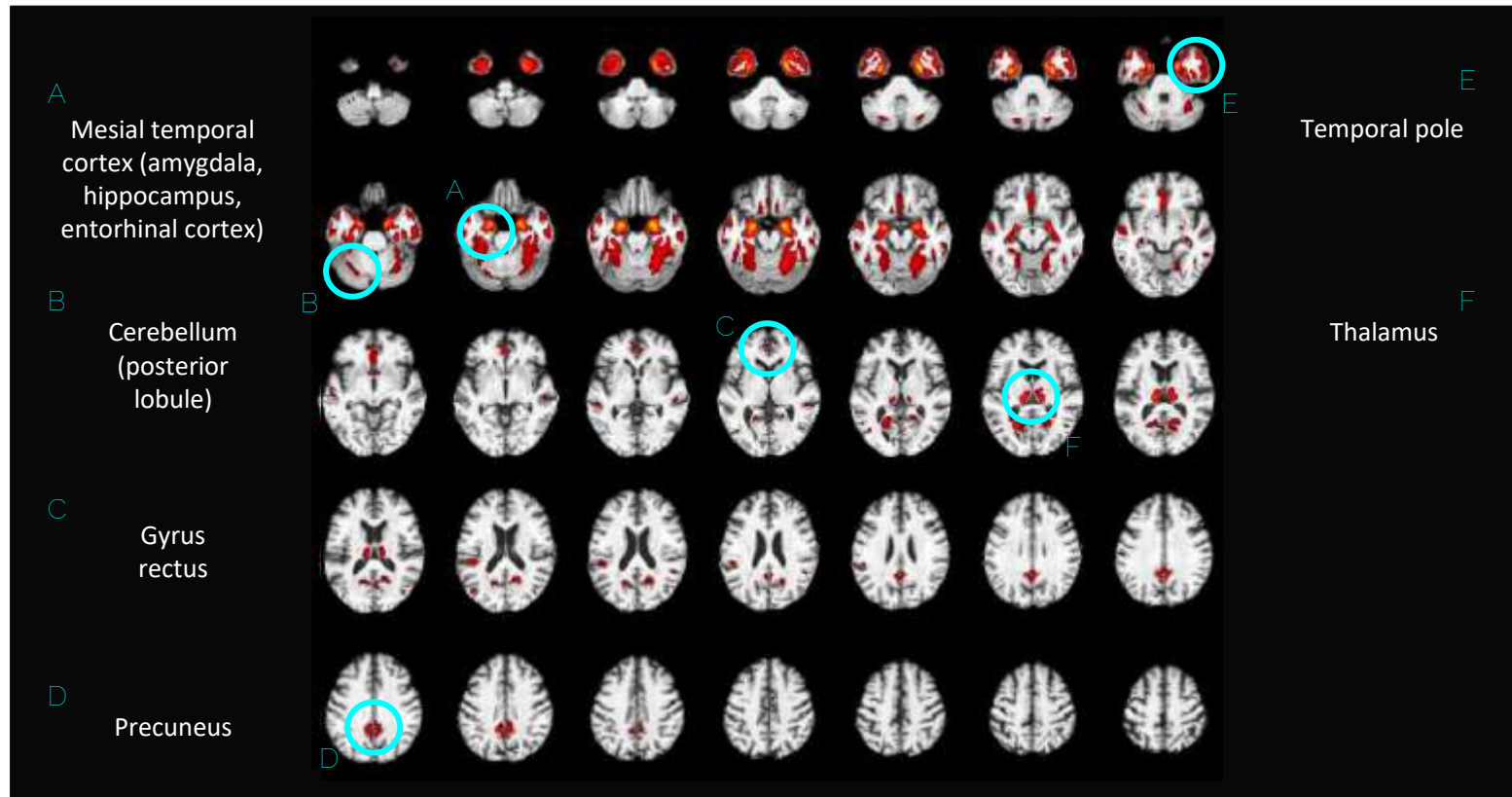


**Best
Structural-MRI
Predictors**

**24 months
before
stable
diagnosis**

MRI characterizes the progressive course of AD and predicts conversion to Alzheimer's dementia twenty-four months before probable diagnosis. *Salvatore et al. 2018, Frontiers in Aging Neuroscience.*

ML models | Examples



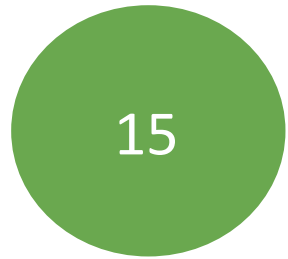
**Best
Structural-MRI
Predictors**

**24 months
before
stable
diagnosis**

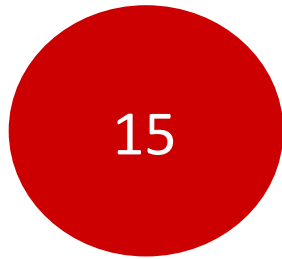
MRI characterizes the progressive course of AD and predicts conversion to Alzheimer's dementia twenty-four months before probable diagnosis. Salvatore et al. 2018, *Frontiers in Aging Neuroscience*.

Autism Spectrum Disorder

Confirm a motor signature of autism



TD



ASD



A dataset of 30
pre-school children (~3 years old)

17 kinematic features
collected during a reach-to-drop task



ASD
TD

Autism Spectrum Disorder
Typically-developing children

Use of machine learning to identify children with autism and their motor abnormalities. *Crippa, Salvatore et al. 2015, Journal of Autism and Developmental Disorders.*



Fig. 1 The experimental task consisted of grasping a rubber ball (2) that was placed over a support (see 1, a); that is, a reach-to-grasp movement before they dropped it in a hole (3). The hole (1, c) was located inside a see-through *square box* (21 cm high, 20 cm wide) and was large enough not to require fine movements. The goal area is

transparent to allow seeing through. 4 markers are placed on the basket under the goal area, 2 on the ball and 3 on each hand (attached to the ulnar and radial surfaces of the participant's wrist and to the hand dorsum on the 4th and 5th metacarpals)

sub-movement 1

the movement necessary to reach the ball and place it on its support

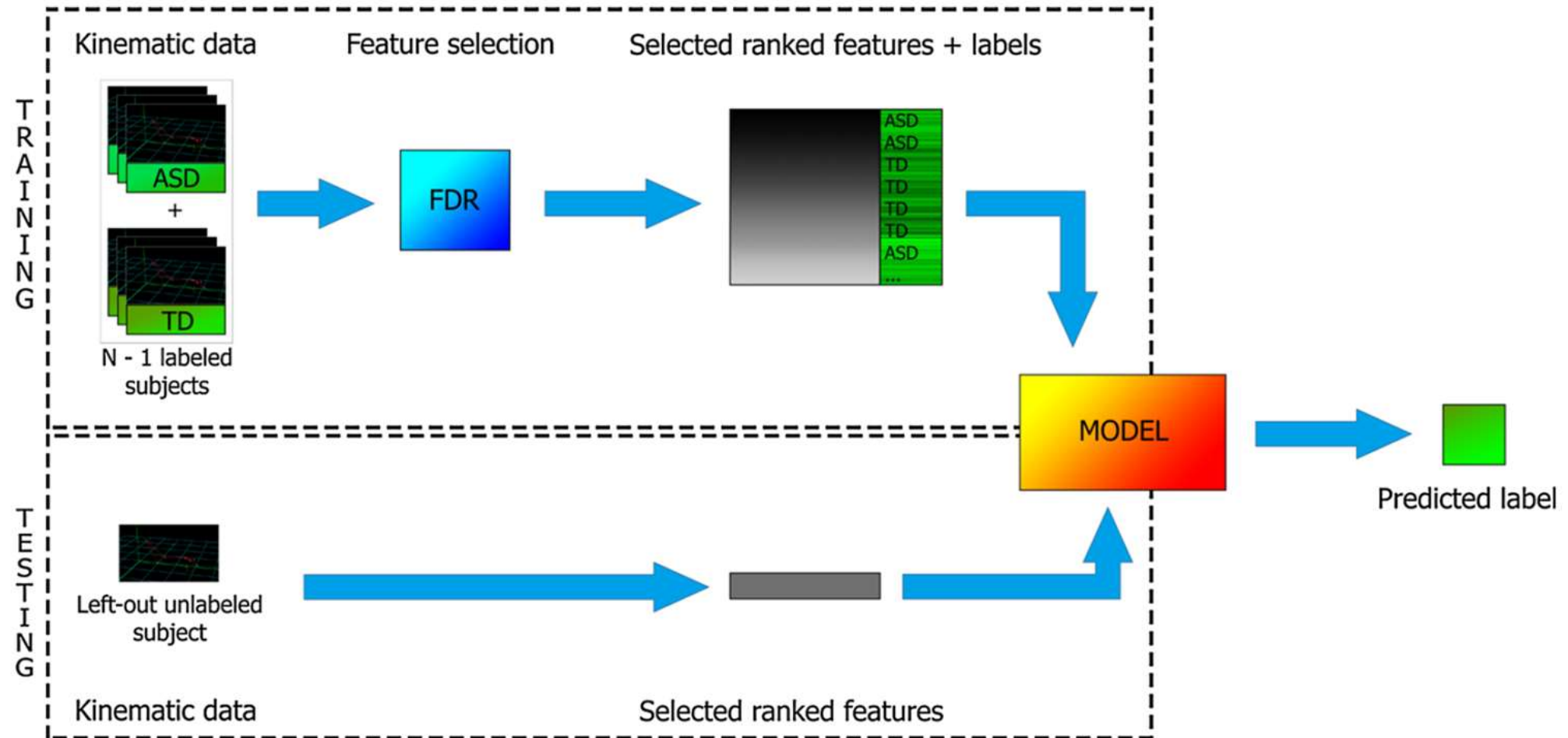
- # movement units
- total movement duration
- peak velocity
- peak acceleration
- time of peak acceleration
- peak deceleration
- time of peak deceleration

sub-movement 2

the movement to transport the ball from a support to the target hole

- # movement units
- total movement duration
- peak velocity
- peak acceleration
- time of peak acceleration
- peak deceleration
- time of peak deceleration
- wrist angle

ML models | Examples



Use of machine learning to identify children with autism and their motor abnormalities. Crippa, Salvatore et al. 2015, *Journal of Autism and Developmental Disorders*.

ML models | Examples

Diagnostic accuracy
(sensitivity / specificity)

Overall
mean

85
(82 / 89)%

Optimal
configuration

97
(100 / 94)%

Use of machine learning to identify children with autism and their motor abnormalities. *Crippa, Salvatore et al. 2015, Journal of Autism and Developmental Disorders.*

Diagnostic accuracy
(sensitivity / specificity)

Overall
mean

85
(82 / 89)%

Optimal
configuration

97
(100 / 94)%

7 optimal features out of 17

sub-movement 2

the movement to transport the ball from a support to the target hole in which the ball was to be dropped

1. total movement duration
2. delta wrist angle
3. # movement units
4. time of peak deceleration
5. peak acceleration
6. time of peak velocity
7. peak velocity

Attention-Deficit/Hyperactivity Disorder

Diagnosis of ADHD & Identification of a signature



TD



ADHD



A dataset of 44
school-aged children (~11 years old)

Multidomain profile of measures

- . biological data
- . neuropsychological data
- . Near-InfraRed Spectroscopy (NIRS)

ADHD
TD

Attention Deficit / Hyperactivity Disorder
Typically-developing children

The Utility of a computerized algorithm Based on a Multi-Domain Profile of Measures for the Diagnosis of attention deficit/hyperactivity Disorder. *Crippa, Salvatore et al. 2017, Frontiers in Psychiatry.*

Multi-domain profile of measures



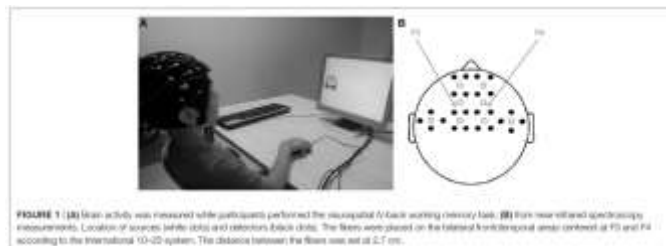
Biological data

10 features



Neuropsychological data

18 features

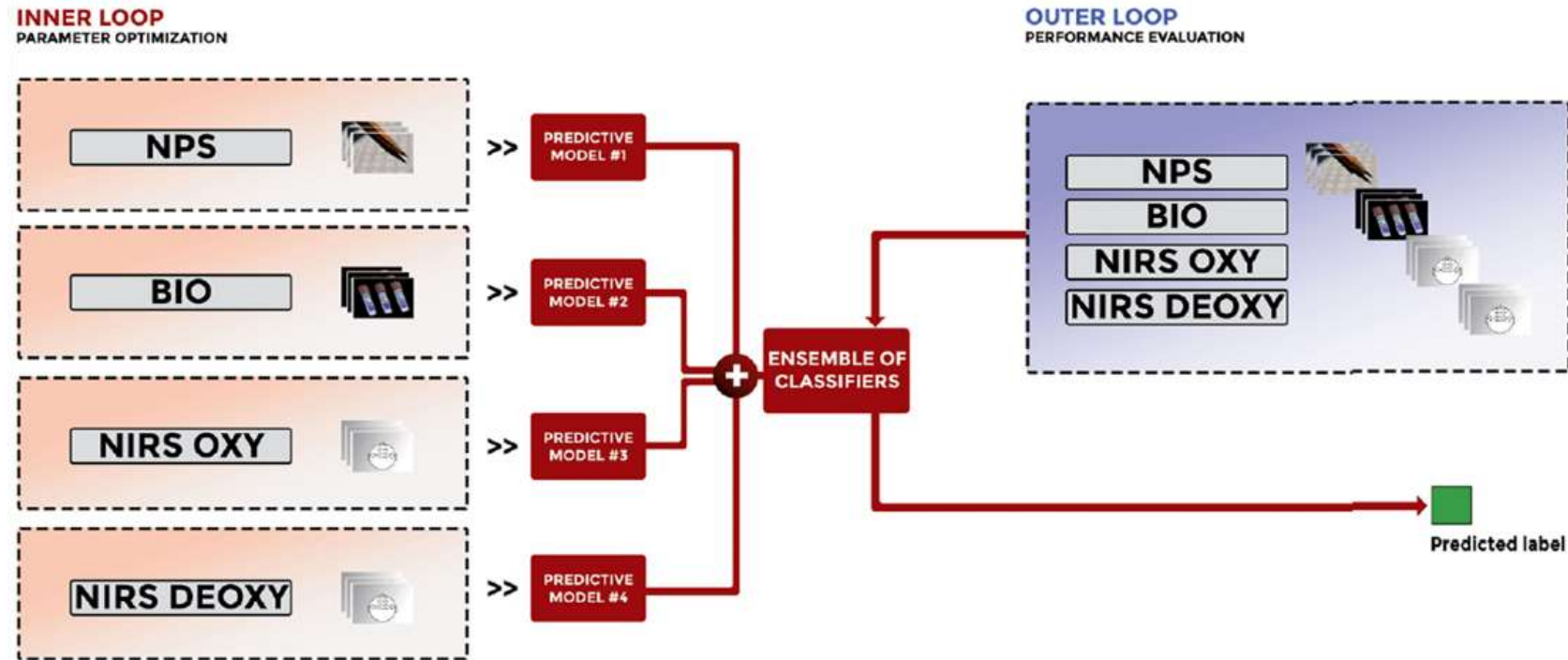


Near-InfraRed Spectroscopy (NIRS)

Oxy/Deoxy data from 32 channels

The Utility of a computerized algorithm Based on a Multi-Domain Profile of Measures for the Diagnosis of attention deficit/hyperactivity Disorder. *Crippa, Salvatore et al. 2017, Frontiers in Psychiatry.*

ENSEMBLE OF CLASSIFIERS



The Utility of a computerized algorithm Based on a Multi-Domain Profile of Measures for the Diagnosis of attention deficit/hyperactivity Disorder. Crippa, Salvatore et al. 2017, *Frontiers in Psychiatry*.

ML models | Examples

Measures	Accuracy (mean ± sd)	Sensitivity (mean ± sd)	Specificity (mean ± sd)
Neuropsychological	62 ± 17	70 ± 27	57 ± 24
Biological	66 ± 21	58 ± 40	73 ± 29
NIRS OXY	57 ± 27	48 ± 47	67 ± 33
NIRS DEOXY	78 ± 22	72 ± 34	82 ± 24
NIRS OXY + DEOXY	72 ± 32	73 ± 29	68 ± 43

biological features

.linoleic acid
.PUFA
.AA
.EPA
.omega-3 index
.AA/DHA
.AA/EPA
.MUFA

linoleic acid and total amount of
polyunsaturated fatty acids

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neuropsychological features

- .sustained attention**-false alarms
- .visual set**-shifting-RT inhibition
- .sustained attention-coefficient of variation
- .visual set-shifting-number of inhibition errors
- .focused attention**-RT correct responses
- .focused attention-correct rejections target non-relevant position
- .focused attention-SD of correct responses RT
- .focused attention-misses
- ...

measures of vigilance, focused and sustained attention, and cognitive flexibility

The Utility of a computerized algorithm Based on a Multi-Domain Profile of Measures for the Diagnosis of attention deficit/hyperactivity Disorder. Crippa, Salvatore et al. 2017, *Frontiers in Psychiatry*.

ML models | Examples

Single domain

Measures	Accuracy (mean ± sd)	Sensitivity (mean ± sd)	Specificity (mean ± sd)
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NIRS DEOXY	78 ± 22	72 ± 34	82 ± 24
NIRS OXY + DEOXY	72 ± 32	73 ± 29	68 ± 43

Multi-domain

Measures	Accuracy (mean ± sd)	Sensitivity (mean ± sd)	Specificity (mean ± sd)
NPS + BIO + NIRS OXY			
NPS + BIO + NIRS DEOXY			
NPS + NIRS OXY + NIRS DEOXY			
BIO + NIRS OXY + NIRS DEOXY			
NPS + BIO + NIRS OXY + NIRS DEOXY			

The Utility of a computerized algorithm Based on a Multi-Domain Profile of Measures for the Diagnosis of attention deficit/hyperactivity Disorder. *Crippa, Salvatore et al. 2017, Frontiers in Psychiatry.*

ML models | Examples

Single domain

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NIRS OXY	57 ± 27	48 ± 47	67 ± 33
NIRS DEOXY	78 ± 22	72 ± 34	82 ± 24
NIRS OXY + DEOXY	72 ± 32	73 ± 29	68 ± 43

Multi-domain

Measures	Accuracy (mean ± sd)	Sensitivity (mean ± sd)	Specificity (mean ± sd)
NPS + BIO + NIRS OXY	71 ± 10	70 ± 27	73 ± 24
NPS + BIO + NIRS DEOXY	81 ± 15	73 ± 24	87 ± 22
NPS + NIRS OXY + NIRS DEOXY	78 ± 18	70 ± 36	87 ± 22
BIO + NIRS OXY + NIRS DEOXY	77 ± 21	63 ± 31	90 ± 21
NPS + BIO + NIRS OXY + NIRS DEOXY	76 ± 16	83 ± 22	68 ± 23

The Utility of a computerized algorithm Based on a Multi-Domain Profile of Measures for the Diagnosis of attention deficit/hyperactivity Disorder. *Crippa, Salvatore et al. 2017, Frontiers in Psychiatry.*