FACULDADE DE ENGENHARIA DA UNIVERSIDADE DO PORTO

Machine learning in the diagnosis of disorders of consciousness using SMART

Manuel Curral



Mestrado Integrado em Engenharia Informática e Computação

Supervisor: Professor Doutor Jaime Cardoso

September 26, 2021

Machine learning in the diagnosis of disorders of consciousness using SMART

Manuel Curral

Mestrado Integrado em Engenharia Informática e Computação

Abstract

People with disorders of consciousness (DOC) are vulnerable to misdiagnosis which can negatively affect their rehabilitation process. The incorrect diagnosis of people with DOC is common, reaching 43 %. This has possible implications for decisions regarding the provision of health care to this group. Diagnosis of this population depends on the assessment of their behavioral responses to stimuli. The intentionality and types of behavior exhibited by people in a vegetative state (VS) and a minimally conscious state (MCS) can be difficult to distinguish and subtle signs of consciousness can go unnoticed. It is widely recognized that the use of standardized and sensitive behavioral assessment scales, such as the Sensory Modality Assessment and Rehabilitation Technique (SMART), can help healthcare professionals to identify subtle signs of awareness. SMART is an assessment tool that combines communication, motor and sensory modalities to diagnose the condition of patients who have suffered severe brain injuries. This method is quite credible and accepted by the healthcare community that deals with this clinical population. It requires and consumes many resources in making the diagnosis.

However, less experienced SMART assessors can be misled by some types of patient responses and even in the analysis of session data, namely in different diagnostic limit zones. Hence a second opinion based on Machine Learning can prove to be very useful. In addition, diagnosis is made session by session and the cumulative diagnostic certainty as the sessions progress. This tool can be detected as well as the expert assessors, in the future it can be very useful to detect in advance (in a smaller number of sessions) the state of consciousness of the patient to be analyzed.

SMART evaluation has already been explored with statistical software such as statistical package for the social sciences (SPSS) combining analysis methods and techniques such as analysis of variance (ANOVA) etc So far, no machine learning methods have been found in partnership with this technique and diagnostic tools (SMART).

The best diagnosis, through a second opinion performed by the machine, is expected to increase the confidence level in decision-making by SMART assessors. More protected and less subject to criticism of negligence, data that possible errors if detected, can be bridged and safeguarded or at least become noticeable therefore, it leads to higher hit rates. Minimizing the allocation of time to human resources for this specific task, can be beneficial for these professionals due to the useful / effective time to perform tasks (elimination of extra hours to do a task that was previously done). Institutions: speeds up the prognosis and diagnosis process, making it financially convenient to make professionals more available, resulting in greater performance and efficiency, with the possibility of performing other essential tasks.

Keywords: Machine Learning, disorders of consciousness, diagnosis, SMART, minimally conscious state, vegetative state, brain injury,

Resumo

Pessoas com perturbações de consciência (PdC) são vulneráveis a diagnósticos errados que podem afetar negativamente o seu processo de reabilitação. O diagnóstico incorreto de pessoas com PdC é comum, podendo atingir os 43%. Isto acarreta possíveis implicações nas decisões relacionadas com a prestação de cuidados de saúde desta população. o diagnóstico desta população depende da avaliação das suas respostas comportamentais a estimulação. A intencionalidade e os tipos de comportamentos exibidos por pessoas em estado vegetativo e estado de consciência mínima, podem ser difíceis de distinguir e sinais subtis de consciência podem passar despercebidos. É amplamente reconhecido que o uso de escalas de avaliação comportamental padronizadas e sensíveis, tal como o Sensory Modality Assessment and Rehabilitation Technique (SMART), pode ajudar os profissionais de saúde a identificar sinais subtis de consciência. O SMART é um instrumento de avaliação que combina funções comunicacionais, motoras e de aferição de sentidos para diagnosticar o estado de pacientes que sofreram lesões cerebrais graves. Este método é bastante credível e aceite pela comunidade da área da saúde que lida com esta população clínica. Ele requer e consome muitos recursos na elaboração do diagnóstico.

Contudo, avaliadores SMART menos experientes podem ser induzidos em erro por algum tipos de respostas dos pacientes e mesmo na análise dos dados das sessões nomeadamente em zonas limite de diagnósticos distintos. Daí uma segunda opinião com base em Machine Learning pode vir a ser muito útil. Além disso, o diagnóstico é feito sessão a sessão sendo a certeza de diagnóstico cumulativa no progredir das sessões.

Esta ferramenta se detetar tão bem como os avaliadores expert, pode no futuro ser muito útil a detetar antecipadamente (num menor número de sessões) o estado de consciência do paciente analisado. - A avaliação SMART já foi explorada através de software estatístico como SPSS usando métodos e técnicas de análise ANOVA etc Até ao momento não foram encontrados métodos de machine learning em parceria desta técnica e ferramenta de diagnóstico (SMART).

melhor diagnóstico, através de uma segunda opinião realizada pela máquina é expectável que aumente o índice de confiança na sua tomada de decisão por parte dos avaliadores SMART. Mais resguardados e menos sujeitos a críticas de negligência, dados que possíveis erros se detectados, possam ser colmatados e salvaguardados ou pelos menos se tornem perceptíveis portanto, conduz a taxas de acerto mais elevadas. Redução dos tempos de alocação a recursos humanos destinados a esta tarefa específica, pode ser benéfico para estes profissionais pelo tempo útil de realização de tarefas (eliminação das horas a mais a fazer uma tarefa já realizada outrora). Instituições: agiliza o processo de prognóstico e diagnóstico, financeiramente conveniente tornar profissionais mais libertos consequente maior rendimento e eficácia com possibilidade de realização de outras tarefas primordiais.

Keywords: Aprendizado Máquina, Perturbações de consciência PdC, diagnóstico de consciência, SMART, estado de consciência mínimo, estado vegetativo, lesão cerebral

Acknowledgements

Hello world!

I want to thank to FEUP to the fantastic conditions to labour, the people who work here and people (students like me) by promoting activities that enable the development of areas of our interest. Dedicate to the people wants to learn and people with a desire to teach in general. It moves the world.

Above all, I would like to thank the fantastic synergies of the Group Coordinator (VCMI) and supervisor Prof. Jaime Cardoso as well the Institutional Coordinator of the Superior School of Health of Porto in name of Prof. Nuno Rocha and Prof. Liliana Teixeira of Polytechnic of Leria as the promoters of this interesting idea.

I want to thank humanity or friendship, the exchange and debate of ideas that exist in the world, which somehow inspires us, gives us encouragement in the most difficult moments and decisions and, on the other hand, gives greater meaning to life, shares happiness and other important feelings with us and vice versa. The power that positive influence can have in one another's lives. The intrinsic willpower that makes us think, research, understand, solve problems and also make us ask for help and listen. My thanks to family, affective, academic/professional, community, financial, sports and health support in general, without you it was not the same thing.

Manuel Curral

"You can't stop the waves, but you can learn to surf."
Jon Kabat-Zinn
"O Universo criou um cérebro para permitir ver-se a si mesmo, para estar consciente de si mesmo."
Henry Markram

Contents

I	Intr	oduction	1
	1.1	SMART	2
	1.2	Objectives	3
	1.3	Structure of the document	3
2	Diag	gnostic tools using Machine Leaning	5
	2.1	Diagnostic tools	5
	2.2	General idea of application	7
3	Mac	chine Learning, a field of AI	9
	3.1	History Perspective	9
		3.1.1 The evolution of AI in board games	11
		3.1.2 Perceptron	11
	3.2	Machine Learning types	11
	3.3	Model Evaluation	16
	3.4	Pipeline steps of a project	16
	3.5	Evaluation process	17
4	Exp	erimental Study	19
	4.1	SMART database considerations	19
5	Con	clusions and Future Work	21
	5.1	Data exploration and analysis	21
	5.2	Machine Learning	21
	5.3	Results	22
	5.4	Further Work	22
Δ	Glos	ssarv	25

X CONTENTS

List of Figures

2.1	MMN topography in patients with disorders of consciousness and in healthy con-					
	trols. The figure shows a comparison of (A) t test maps from Boly et al. (1) for					
	the MMN (comparison of deviant and standard trials) with (B) similar maps ob-					
	tained from 120 recordings collected in the past three years in Lionel Naccache's					
	laboratory, Hôpital de la Salpêtrière, Paris					
3.1	Machine learning and directly related areas					
3.2	Types of AI					
3.3	Perceptron model representation					
3.4	Human evaluation vs different types of AI paradigms					
3.5	Visual Explanation of SVM					
3.6	Visual Explanation of the kernel trick					
3.7	Taxonomy of supervised learning algorithms					
3.8	Main features of the algorithm					
3.9	Confusion Matrix					
3.10	full matrix confusion table with metrics included					
5.1	Confusion Matrix and ROC curve of a Bunch of Algorithms					
5.2	Neural Network Confusion Matrix and ROC curve resulting from the iterative					
	method with 100 epochs, 25 neurons and ADAM optimizer					

xii LIST OF FIGURES

List of Tables

1.1		1
4.1	Classification with 2 states	20
4.2	Classification with 3 state	20
4.3	Describe Age and Time	20
4.4	Gender code definition	20
4.5	Etiology code definition	20

xiv LIST OF TABLES

Abbreviations

API Application Programming Interface

ANOVA Analyis of Variance

DOC Disorders of Consciousness
 DRS Disability Rating Scale
 GCS Glasgow Coma Scale
 ICC Intra Class Correlation

fMRI functional Magnetic Resonance Imaging

MCS- Minimally Conscious State Minus MCS+ Minimally Conscious State Plus MMN Multifocal Motor Neuropathy

NN Neural Network

SMART Sensory Modality Assessment and Rehabilitation Technique

WNSSP Western Neuro Sensory Stimulation Profile

TBI Traumatic brain injury

VS Vegetative State WWW World Wide Web

Chapter 1

Introduction

Context and Motivation

Consciousness is the individual's ability to be aware of the knowledge of self and the environment. Furthermore, the ability to respond to various voluntary internal and external stimuli [1]. In basic neurological terms, it is composed of awareness and wakefulness [2]. The different states of consciousness are represented in the table [3].

Condition	Wakefulness	Awareness
Coma	-	-
Vegetative State	+ to ++	-
Minimally Conscious State	+ to ++	+
Emerged from Minimally Conscious State	++	++

Table 1.1

Brain Injury (BI) is head injury that damage in the brain and his complex connections.

This causes problems with how a person can think and interact with the world around him or her. Following a brain injury, there are specific cognitive skills that are no longer functioning in the same capacity [4]. Perception, observation, and recognition of information are deeply affected.

There are events that damage areas of the brain that control parts of the human body. And the patient's faculties are conditioned. The origin of brain injuries can be:

- acute: as in a virus or haemorrhage
- traumatic: like road accidents, impacts where there are head injuries
- non-traumatic: such as drowning, sudden attacks on organs and consumption of substances harmful to the body

That cause brain damage that leads to consequences in terms of disturbing the person's consciousness [5].

2 Introduction

After coma, the rates of diagnostic errors, namely in the distinction between vegetative state (VS) and the minimally conscious state (MCS), are high $\approx 40\%$ [6] [7] [8].

The diagnosis of the condition of the patient who suffers brain damage is made using scales to assess the behavioural response to stimulation: CRS-r, SMART, WHIM, WNSSP, Rancho levels physicians etc [9]. With technological advancement and the spread of its use, neuronal imaging technologies such as: functional magnetic resonance imaging (fMRI), electroencephalogram (EEG) positron emission tomography (PET) [10] [11].

It has been possible to leverage the discovery and knowledge in the area of neuroscience and the identification, consequent diagnosis, of the cerebral behaviour of patients [4]. This helps a lot in studying DOC because it demonstrates the level of responsiveness that could not be obtained by the behaviour scales already mentioned 1.

VS is defined as the absence of self-awareness and the environment 1.1. Behaviours are limited to reflexive activities indicating no purposeful movement, neither experience of suffering or evidence of comprehension [12].

MCS is serious but does not represent a complete lack of awareness resulting from widespread damage to the telencephalon (the part of the brain that controls thinking and behaviour). The intentionally and types of behaviour exhibited by people in a VS and a MCS, can be challenging to distinguish, and subtle signs of consciousness can go unnoticed. It is widely recognized that the use of standardized and sensitive behavioural assessment scales, such as the Sensory Modality Assessment and Rehabilitation Technique (SMART), can help healthcare professionals identify subtle signs of awareness.

With the data gathered from various patient assessments, we think it is possible to train an artificial intelligence model. Able to recognize hidden patterns of human knowledge or minute details resulting from the multiple data that feed the model. In order for the model to generalize the upcoming new data and produce a diagnosis accurate enough to be of help to SMART evaluators [13].

1.1 SMART

SMART is an assessment tool that combines communication, motor and sensory modalities to diagnose patients who have suffered severe brain injuries [14].

Five levels: no response, reflex response, withdrawal response, localized response and differentiated response.

The main advantage of the method is entirely credible and accepted by the healthcare community that deals with this clinical population.

The disadvantage is that it requires and consumes many resources in making the diagnosis.

Research continues on clinical tools such as (fMRI) with improved diagnostic certainty and prognostic applications [4]. There are 3 main factors that influence the prognosis of patients in the Vegetative State (VS) and the patient's minimum state of consciousness (MCS):

1.2 Objectives 3

• Time (the longer you stay in the state, the more complicated functional recovery becomes)

- Age (young people have a higher recovery rate, linked to physiological recovery processes and brain plasticity)
- Type (if non-traumatic, there is a shorter potential recovery window)
- Note: The more severe the degree of injury, the rarer the recovery.

Since the diagnostic evaluation is done over a period of 3 weeks over 10 sessions, distributed in equal numbers in the morning and afternoon.[9]

1.2 Objectives

The goal of this dissertation is to give greater certainty in the diagnosis, or for instance reduce levels of misdiagnosis of disorders of consciousness. The following steps materialize the aim:

- 1. Classification in 2 possible stages (minimal state of consciousness and vegetative state)
- 2. Reduce time procedures diagnosis: after the first complete goal check, reduce the number of sessions to less than 10
- 3. Try to find correlations between origin and possible stages and gauge the accuracy of results with basis on number of sessions available

1.3 Structure of the document

In total, the dissertation report has 5 chapters. T Chapter with state of the art 2.

Chapter of Machine Learning, a field of AI 3.

Evaluation Metrics Chapter 4.

Conclusions Chapter 5.

Areas:

 $CCS \rightarrow Computing$ methodologies $\rightarrow Machine$ learning

CCS →Applied computing →Life and medical sciences

4 Introduction

Chapter 2

Diagnostic tools using Machine Leaning

2.1 Diagnostic tools

Neuroscientific technologies, experimental methods study the patients' brain processes through many devices, in specific:

Neuroimaging, psychophysical techniques or psychological tests are used to study processes such as learning, attention, memory, or emotion [15].

The following stand out:

• Electroencephalogram (EEG)

Electroencephalography is a simple, non-invasive technique based on the recording and evaluating brain activity using electrodes placed on the skull surface[16] [11]. The (EEG) is the record that results from the measurement of the electrical potentials of the brain [4] [17]. EEG shows the electrical fluctuation in the different locations of the cortex [18]. However, it has the disadvantage of having an insufficient resolution to register the neuronal activity in deeper brain structures, such as the *nucleus accumbens* related to the processing of emotions[19].

Study using MMN tomography technique resulting in classification of patients with DOC in figure 2.1

• (Functional) Magnetic Resonance

It is a method of diagnosis and fundamental research in Market research analysing emotions planning surgery brain mapping neurosciences [21]. It allows the analysis of the subjects' response to different activities or stimuli. Magnetic resonance imaging is a non-invasive technique used to obtain information about the subjects' response to different stimuli (very common in research and analysis of neurological diseases such as Alzheimer's Disease and

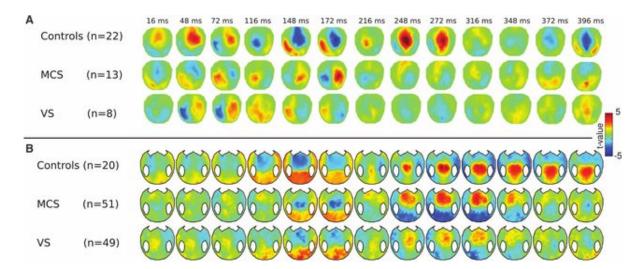


Figure 2.1: MMN topography in patients with disorders of consciousness and in healthy controls [10] [20]

also for soft tissue injuries and inflammation). This technique evaluates the activation and the emotional state of the subject when exposed to certain stimuli [22].

• GSR (galvanic skin response)

Measurement of the galvanic response is done by placing electrodes on the fingers. Studies measure resistance skin and its conductance.

A GSR amplifier applies constant tension to the skin, of such a low voltage that the individual cannot perceive it through electrodes. The current generated in the skin by tension can be detected and recorded. The output of the GSR amplifier determines the conductance. The conductance of the skin gives feedback on the body's response to the stimulus, it is widely used in post-coma and coma phases [23, 24].

· Eye-tracking

Eye-tracking refers to recording the movements of an individual's eye while examining a visual stimulus. In broad, it is responsible for measuring eye movements using a camera that quantifies them. Modern eye trackers record eye position and movement using contrast to locate the central point of the pupil and create a reflection of the cornea using infrared light. It is possible to perform the analysis of the position of the gaze and the movement of the eyes in three-dimensional environments [25]. Eye-tracking techniques apply to domains where interaction and interests matter, looking to sell or immerse to improve the customer or user experience. In medicine, it is essential to recognize behaviours and patterns to investigate further and be able to characterize [26].

• Face reading

Facial expressions are one of the most robust visual methods for conveying emotions. The face plays a crucial role in the cognitive processes of individuals since the signs that show

facial expressions denote internal states or emotions. The analysis of facial expressions provides valuable information when combined with other tools that allow sensory information collection, such as eye-tracking or EEG.

2.2 General idea of application

In medicine and biomedical (health in general) areas, it is common, there are pre-determined datasets for training and a new dataset(s) for testing. Typical general use case: Models based on the knowledge of n patients, trying to classify the status of other patients who will arrive [27].

The increase in the processing and computing power of the systems, high performance computing (which includes GPU's, CPU's, high-speed interconnect, memory, libraries, storage, compilers etc), allowed algorithms that would take months to be used in good time using even non-own resources, to distance, as in the case of cloud computing A [28].

Chapter 3

Machine Learning, a field of AI

Machine Learning is a subject field of Artificial Intelligence. Artificial Intelligence (AI) is an expansive branch of computer science whose fundamental aim is to develop computational systems (smart machines) more useful and capable of performing tasks that typically require human intelligence, showing operational behaviours [29].

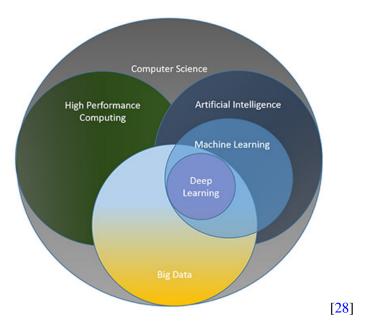


Figure 3.1: Machine learning and directly related areas

3.1 History Perspective

The field of AI began in the 1940s with Alan Turing, an English mathematician and computer scientist. He is considered as father of Artificial Intelligence and Computer Science [30]. This outstanding achievements include:

- Cracking the Enigma code which, according to Keegan, was allegedly decisive in the course of the Second World War shortening it and contributing to this known outcome [31].
- Inventor of first models of the general purpose computers: Turing machines [32]
- Creator of a Turing test, currently used, to find out whether an intelligent agent is capable of thinking like a human being. Through a conversation in natural language between a human and a computer, a human evaluator have to try to distinguish the computer from the human. When this happens, the test is negative. Else, it means that computational system achieve a human like level of artificial intelligence [33].

Until now applications have been developed in the Narrow AI branch, this means that AI tries to perform well on one task, rather than strive for human intelligence, named Artificial General Intelligence (AGI) or beyond the best human brains of that, named Artificial Super Intelligence (ASI).

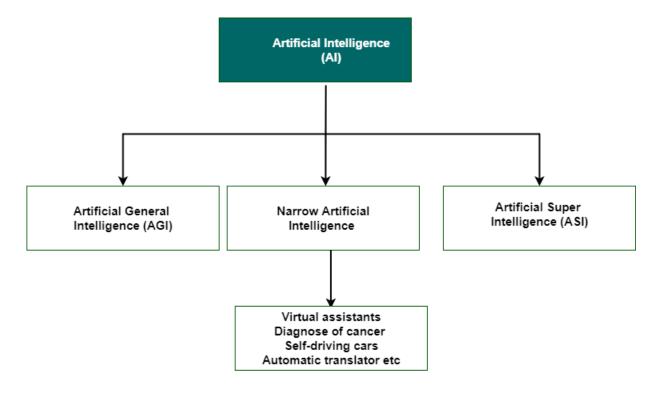


Figure 3.2: Types of AI

To reach the same objectives that make people seem intelligent behind the study of those processes/principle that make computers to execute tasks that are similar to those of the humans in stereotyped situations and more effective way [34]. Techniques of programming used, like non-deterministic search, are based, at least partially, on declarative languages, namely logic-based, functional or at least, object-oriented.

3.1.1 The evolution of AI in board games

Theory of neural network (NN) begins in 1943 using electrical circuits as a nerve cell with a logic gate with binary outputs. In the 1950s, computer scientists simulate the idea in their work, a hypothetical NN.

Over time, we have seen successive application of artificial intelligence in board games, because they are both understandable and complex.

Followed events chronologically:

- In the 50's, the investigation on the game of checkers stands out. So learning by playing and giving a set of rules game as well a sense of direction and some parameters with not including the weight of your importance. Arthur Samuel developed software that helped the IBM computer get better at checkers, the more it played. With the premise of the possibility of scaling and transposing these machine learning techniques to other areas [35].
- Already in 1997, IBM's Deep Blue (chess-playing computer), capable of evaluating 200
 million positions per second (twice as fast as the 1996 version), beat grandmaster Garry
 Kasparov.
- And more recently in 2016, human intelligence head to head with machines, AlphaGo AI program powered by Google. In the game Go, considered as the oldest in the world and probably the most complex. Google DeepMinds AlphaGo combines reinforcement learning techniques with simulation so is powered with all types of Go matches, and then study, learn from them, and discover your own moves. It knows the rules, and discovers highly efficient creative moves never before thought by humans [36].

Due to the evolution of the processing power of computers, it is now possible to run computationally demanding algorithms in terms of calculations and/or the amount of data used [37]. In this way, there was a path that made possible a new era, a revolution of Artificial intelligence called deep learning, that mimics neural networks of the human brain.

3.1.2 Perceptron

The Perceptron is like an atom of artificial intelligence that performs computations to extract information or knowledge from input data [38].

3.2 Machine Learning types

Machine learning algorithms are categorized into three main categories:

- supervised learning 3.2.0.1
- · unsupervised learning
- · reinforcement learning

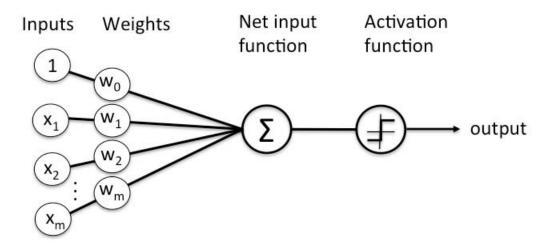


Figure 3.3: Perceptron model representation [39]

3.2.0.1 Supervised learning

The process of creating the machine learning model involves having a dataset that contains the features and labels in order to use it to classify any given new point of data.

Within supervised learning we can divide into:

- Classification performing as a category the class or label (y)
- Regression if we consider as a class(y) a continuous variable

Examples:

Linear Regression, assumption: the class is expected to be a linear combination 3.2.0.1 of the features! It could be adapted for higher dimensions.

$$f(X) = \beta_0 + \beta_1 X_1 + \beta_2 X_2 + \dots + \beta_p X_p$$

There are two types of algorithms:

- Lazy like:
 - K-Nearest Neighbours or KNN (distance basic technique)
 When a new point appears, we calculate the similarity (or distances, as in clustering) of the K-nearest Neighbours. Where k is a positive number, to obtain optimal value should use an error plot or accuracy plot. The class of our new input is assigned to the class most common, like voting system in democracy [27].

- Rule Based

A technique widely used in healthcare in the 70s. However, it has its limitations in comparison with the techniques used today. Those limitations include high monetary cost, dependence on specialized human resources, and complexity of how to update the rules. Practical implementation constraints are illustrated in the table below.

Approaches	Model comprehensibility	Performance	Reproducibility	Dependency on prior knowledge	Development and training costs ^a	Running costs	Around- the-clock availability	Update costs
Human evaluation	High	Moderate or high	Moderate	High	High	High	Low	High
Rule-based algorithms	High	Moderate or high	High	High	Moderate or high	Low	High	High
Feature-based machine- learning methods	Moderate or high	Moderate or high	High	Moderate ^b	Moderate	Low	High	Moderate ^c
Deep artificial neural networks	Low or moderate	High	High	Low	Moderate	Low	High	Low

*The estimated cost of training professionals that carry out the clinical tasks (human evaluation) or of developing the automated system (rule-based, feature-based or deep-artificial-neural-network-based) that performs the tasks. For feature-based machine-learning methods, prior knowledge may facilitate the derivation of useful features from the raw data. When the update requires encoding new features, the update cost of feature-based machine-learning methods includes feature engineering and model retraining.

Figure 3.4: Human evaluation vs different types of AI paradigms.

[13]

 Naive Bayes Rules its name comes from the Bayes theorem on which it is based, applies and calculates conditional probabilities and probabilities. Suitable for large and unknown datasets. Assumes that features are independent or unrelated between with each other. Then the algorithm learns and collects information from each feature class individually.

• Eager like:

- Neural Networks
- SVMs Intends to maximize the decision boundary margin. The orange dots are the support vector, the points closer to the boundary of decision.

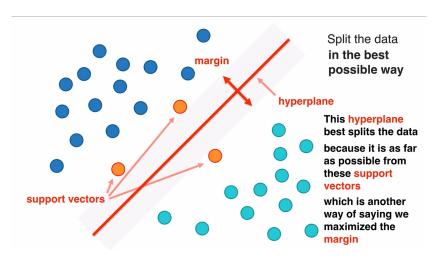


Figure 3.5: Visual Explanation of SVM [40]

When the problem is not linearly separable use the kernel trick:

- * Transition to a higher dimensional space feature space
- * Get an optimal boundary
- * Return to input space

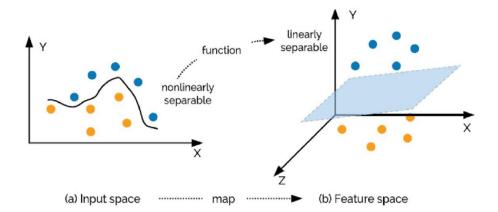


Figure 3.6: Visual Explanation of the kernel trick

- Decision trees

Hierarchical learning (decomposes in subtrees), to build one:

- 1. Select feature for root node and create a branch for each possible feature value
- 2. : split instances into subsets: one for each branch extending from the node
- 3. Repeat for each branch, using only the instances that reach the branch

A selected feature corresponds a question or "test" to the database in order to subdivide data until obtaining a branch of the same class called pure leaves. The goal is to pick the most significant features to subdivide in the faster way possible (minimum number of tests), avoiding overfitting. Note that only 1 feature is picked by the tree if there are more that encodes the same information. The benefits comparing with another algorithms is ease of being understandable by the people, the ability to visualize beyond the algorithms doesn't variant to scaling of the data.

Random Forests

Multiple decision trees (ensemble method), hierarchical learning, divide and conquer method. Bootstarping of samples for each decision tree, and all decision trees, makes a prediction. In regression, makes an average of results. For other side in classification, makes a soft voting that consist that: the most probable class, the result of the average of the probabilities of all the probabilities of the predictions of the decision trees, is chosen with the representative label [41].

- Gradient boosted regression trees:

- * used for regression and classification
- * considered one of the most powerful boosting algorithms and widely used on supervised learning, nevertheless is required careful tuning the parameters.

- * have a strong pre-pruning which results in shallow trees, typically no more than
 5 levels deep consequently is smaller in memory and faster to predict. With more trees and n_estimators allow more chances to correct mistakes on training set.
- * make a serialization of trees with the aim of predicting the error value of the previous tree. And with learning rate parameter, manage the intensity of correctiveness in previous trees to the definition in its final result.

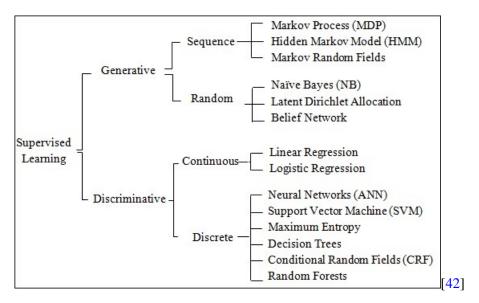


Figure 3.7: Taxonomy of supervised learning algorithms

In this project we focus, use and explore Naive Bayes and a bunch of supervised algorithms of the discriminative and discrete type according to the input data.

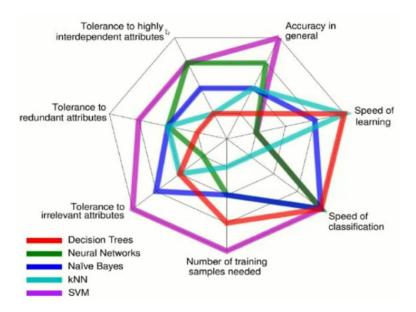


Figure 3.8: Main features of the algorithm

3.3 Model Evaluation

The common practice in the field is to evaluate the model to see if it has the desired performance.

3.4 Pipeline steps of a project

- Collecting the data
- Get some features
 - Extract, Analyse and Select
- Get a model
- Train a model
- Test the model
- Evaluate the model, adjust and repeat if necessary

It is worth pointing out that in case of data limitation, it is important to test with new data. In the evaluation phase, it is expected that:

- A model is able to generalize from seen data to unseen data.
- Is "Able" according to some metrics

Furthemore, there are statistical methods that allow us to assess the model in terms of predictions and get more information.

3.5 Evaluation process

• We split the dataset into a training dataset and a test dataset. Under no circumstances we should use the "test" dataset to train and perform evaluation.

Otherwise, the data will be skewed. That may lead to the misdiagnosis of performance at levels higher than the real ones. Testing the model will take an example from dataset, evaluate it without adjusting the model. The evaluation result or prediction of the model will be compared with the groundtruth labels.

- The groundtruth labels are typically not given to the model to make the prediction.
- They are used to confirm if the prediction was correct or not.

Let us examine a simple example with two categorical classes: "yes" or "no" If a dataset example has a label "yes" and the model predicted as a "yes", the model as correctly classified the example. Otherwise, it is an erroneous prediction.

For each input x with groundtruth class y we have: Correctly discriminated/classified examples Incorrectly discriminated/classified examples

		Predicted Class		
		Positive	Negative	
Class	Positive	True Positives (TP)	False Negatives (FN)	
Actua]	Negative	False Positives (FP)	True Negatives (TN)	

Figure 3.9: Confusion Matrix [43]

From the data in the other table, we can deduce a set of metrics [44].

It is good practice to save this data (confusion matrix) because from it, we can calculate everything from these metrics [46].

		Predicted c	ondition		
	Total population = P + N	Positive (PP)	Negative (PN)	Informedness, bookmaker informedness (BM) = TPR + TNR - 1	Prevalence threshold (PT) = √TPR×FPR - FPR TPR - FPR
Actual condition	Positive (P)	True positive (TP),	False negative (FN), type II error, miss, underestimation	True positive rate (TPR), recall, sensitivity (SEN), probability of detection, hit rate, power $= \frac{TP}{F} = 1 - FNR$	False negative rate (FNR), miss rate = $\frac{FN}{P} = 1 - TPR$
	Negative (N)	False positive (FP), type I error, false alarm, overestimation	True negative (TN),	False positive rate (FPR), probability of false alarm, fall-out $= \frac{FP}{N} = 1 - TNR$	True negative rate (TNR), specificity (SPC), selectivity = $\frac{TN}{N} = 1 - FPR$
	Prevalence $= \frac{P}{P+N}$	Positive predictive value (PPV), precision = TP = 1 - FDR	False omission rate (FOR) $= \frac{FN}{PN} = 1 - NPV$	Positive likelihood ratio (LR+) = TPR FPR	Negative likelihood ratio (LR-) = FNR TNR
	Accuracy (ACC) $= \frac{TP + TN}{P + N}$	False discovery rate (FDR) = FP = 1 - PPV	Negative predictive value (NPV) $= \frac{TN}{PN} = 1 - FOR$	Markedness (MK), deltaP (Δ p) = PPV + NPV - 1	Diagnostic odds ratio (DOR) = LR+ LR-
	Balanced accuracy (BA) $= \frac{TPR + TNR}{2}$	$F_1 \text{ score}$ $= \frac{2PPV \times TPR}{PPV + TPR} = \frac{2TP}{2TP + FP + FN}$	Fowlkes–Mallows index (FM) = $\sqrt{PPV \times TPR}$	Matthews correlation coefficient (MCC) = √TPR×TNR×PPV×NPV - √FNR×FPR×FOR×FDR	Threat score (TS), critical success index (CSI),

Figure 3.10: full matrix confusion table with metrics included [45]

Chapter 4

Experimental Study

4.1 SMART database considerations

This project consists of taking up the technique of diagnosing patients with consciousness disorders. For that, I had access to a dataset with 35 records of the evaluation of this technique that took place in a hospital in London, where Professor Liliana Teixeira worked. Professor Liliana allowed me to access this dataset and was kind enough to explain how this technique works and clarify all the doubts that followed from reading the data and its particularities. The dataset has 147 columns of feature data. Many columns are repeated from 10 sessions that the technique uses for the evaluation of 8 modalities. There are 5 sensory modalities (gustatory, olfactory, auditory, visual and tactile), the rest are functional communication, wakefulness/arousal and motor function [47]. In this way, 8 modalities are evaluated in 10 times, making up 80 features that enlarge the data set. The multiple assessments of the same modality are, often, synonymous with feature redundancy. The individual assessment of each session is done at the end of each session and this technique covers 2 evaluators, in 10 sessions and each of these is associated with the degree of certainty. In total we have over 40 columns/features. In addition, there is a subsequent evaluation where the evaluators discuss and reach a consensus on the joint result for each session and the associated degree of certainty. So we have 20 more features. Of the other 7 features that remain, 2 are for patient identification and their evaluation order. These two features introduce noise to the model creation and not taken into consideration while creating supervised learning models. Another 2 columns correspond to the y or diagnostic labels (in case of 2 and 3 possible states). The labelling mode is shown in the following tables:

Table 4.1: Classification with 2 states

Label	Condition	Quantity
1	Vegetative State	10
2	Minimally Conscious State	25

Table 4.2: Classification with 3 state

Label	Condition	Quantity
1	VS	10
2	MCS-	11
3	MCS+	14

The **time** variable is the time interval, in months, between brain injury and SMART assessment. Age refers to the patient's age. These features are continuous, so, in the following table, you can check information about these:

Table 4.3: Describe Age and Time

Features	Mean	Minimum	Maximum	Median
Time (in months)	8,2	3	70,0	52,0
Age (in years)	49,7	19,0	77,0	6,0

The other two features are the patient's gender and the etiology/origin of the disease, they are binary with the following meaning:

Table 4.4: Gender code definition

Label	Gender	Quantity
1	Female	7
2	Male	28

Table 4.5: Etiology code definition

Label	Etiology	Quantity
1	Traumatic	11
2	Not traumatic	24

Chapter 5

Conclusions and Future Work

5.1 Data exploration and analysis

Upon close examination of the data summary we can surmise that the classification for the two states is unbalanced. 4.1.

And it turns out that there are fields with null values, 111 in total. This happens when the patient is blind or the gustatory sensory system is not possible to assess. It happens in the situations like patients are fed through a gastric tube or other alternative feeding mechanisms such as transcheostemia etc All 'NA' are changed and assigned a non-significant value: -1. The reason for doing so is that data is precious and scarce and removing records would further shorten our dataset. If we do not have sufficient Without data you cannot make the algorithm make a perfect model. The id and order insignificant columns are removed. And the column of the diagnosis made by human evaluators, our label, is separated from X for an isolated panda's series y.

5.2 Machine Learning

X and y are split into training and testing subsets. Taking care to stratify the data in relation to the label (stratify=y). This is to ensure that all classes are proportional in relation to training side to the testing side, avoiding the risk of poorly trained classes.

5.3 Results

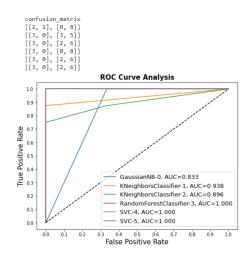


Figure 5.1: Confusion Matrix and ROC curve of a Bunch of Algorithms

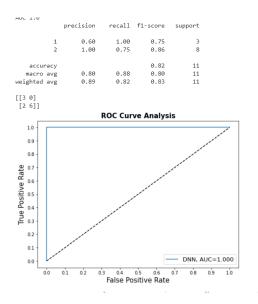


Figure 5.2: Neural Network Confusion Matrix and ROC curve resulting from the iterative method with 100 epochs, 25 neurons and ADAM optimizer

5.4 Further Work

The project can be futher developed to find correlation of features in algorithms. Divide in two subgroups based on etiology. We can also see if it is possible to do less sessions to minimize resources of this tecnique(human resources, time spend). Since it may no longer be adding added value to the data.

5.4 Further Work 23

• A broader view of classification in 3 possible stages (vegetative state, minimal state of consciousness minus and plus)

- Iterate over the various machine learning algorithms
- Unification of similar features [48]
- Predict the correlation of selected features and limit the number of sessions needed
- Use of diagnostic tools to sensory data acquisition
- Test the model with new and real data where it does not contain as rather sessions
- Automate the entire process, acquire technological tools that allow you to track patients regularly

Appendix A

Glossary

Coma: from the Greek kôma = deep sleep, can be defined as a state of total or partial loss of consciousness, voluntary motor skills and sensitivity, generally due to brain damage, intoxications, metabolic and endocrine problems, without which, regardless of severity, vital functions are maintained to a greater or lesser degree (1). When physiological, the state of coma can be measured using the Glasgow Coma Scale (GCSI) and when pharmacological using the Ramsay Sedation Scale (RSS).

Cloud Computing: technology that allows remote use of computing resources through Internet connectivity

26 Glossary

Bibliography

- [1] A. M. Owen, "Disorders of consciousness," *Annals of the New York Academy of Sciences*, vol. 1124, no. 1, pp. 225–238, 2008.
- [2] Z. S. Zheng, N. Reggente, E. Lutkenhoff, A. M. Owen, and M. M. Monti, "Disentangling disorders of consciousness: Insights from diffusion tensor imaging and machine learning," *Human brain mapping*, vol. 38, no. 1, pp. 431–443, 2017.
- [3] Hong Song, Qiang Li, Shixiong Li, Wei Kang, Jian Yang, Yi Yang, and Jianghong He, "Classification of patients with disorder of consciousness based on DTI sequence analysis," 2017 IEEE International Conference on Smart Cloud (SmartCloud). Proceedings, 2017.
- [4] A. Bender, R. J. Jox, E. Grill, A. Straube, and D. Lulé, "Wachkoma und minimaler Bewusstseinszustand: Systematisches Review und Metaanalyse zu diagnostischen Verfahren," *Deutsches Arzteblatt International*, vol. 112, pp. 235–242, apr 2015.
- [5] L. Teixeira, D. Blacker, and N. Rocha, "Disorders of consciousness," *Multidisciplinary Interventions for People with Diverse Needs-A Training Guide for Teachers, Students, and Professionals*, p. 27, 2020.
- [6] K. Andrews, L. Murphy, R. Munday, and C. Littlewood, "Misdiagnosis of the vegetative state: retrospective study in a rehabilitation unit," *Bmj*, vol. 313, no. 7048, pp. 13–16, 1996.
- [7] H. Gill-Thwaites and R. Munday, "The sensory modality assessment and rehabilitation technique (smart): a valid and reliable assessment for vegetative state and minimally conscious state patients," *Brain injury*, vol. 18, no. 12, pp. 1255–1269, 2004.
- [8] C. Schnakers, A. Vanhaudenhuyse, J. Giacino, M. Ventura, M. Boly, S. Majerus, G. Moonen, and S. Laureys, "Diagnostic accuracy of the vegetative and minimally conscious state: clinical consensus versus standardized neurobehavioral assessment," *BMC neurology*, vol. 9, no. 1, pp. 1–5, 2009.
- [9] H. Gill-Thwaites and R. Munday, "The sensory modality assessment and rehabilitation technique (SMART): A valid and reliable assessment for vegetative state and minimally conscious state patients," *Brain Injury*, vol. 18, no. 12, pp. 1255–1269, 2004.

28 BIBLIOGRAPHY

[10] J. R. King, T. Bekinschtein, and S. Dehaene, "Comment on "preserved feedforward but impaired top-down processes in the vegetative state"," dec 2011.

- [11] C. Da and S. Lourenço, "Deep Learning for EEG Analysis in Epilepsy," 2019.
- [12] M.-S. T. F. on PVS, "Medical aspects of the persistent vegetative state," *New England Journal of Medicine*, vol. 330, no. 21, pp. 1499–1508, 1994.
- [13] K.-H. Yu, A. L. Beam, and I. S. Kohane, "Artificial intelligence in healthcare," *Nature Biomedical Engineering 2018 2:10*, vol. 2, pp. 719–731, oct 2018.
- [14] L. da Conceição Teixeira, H. Gill-Thwaites, F. Reynolds, and S. Duport, "Can behavioural observations made during the SMART assessment detect the potential for later emergence from vegetative state?," *Neuropsychological Rehabilitation*, vol. 28, pp. 1340–1349, nov 2018.
- [15] S. Corchs, G. Chioma, R. Dondi, F. Gasparini, S. Manzoni, U. Markowska-Kacznar, G. Mauri, I. Zoppis, and A. Morreale, "Computational methods for resting-state EEG of patients with disorders of consciousness," 2019.
- [16] O. Avilov, S. Rimbert, A. Popov, and L. Bougrain, "Deep Learning Techniques to Improve Intraoperative Awareness Detection from Electroencephalographic Signals," in *Proceedings* of the Annual International Conference of the IEEE Engineering in Medicine and Biology Society, EMBS, vol. 2020-July, pp. 142–145, Institute of Electrical and Electronics Engineers Inc., jul 2020.
- [17] Z. Xu, J. Wang, R. Wang, Z. Zhang, and S. Yang, "Deep Convolutional Neural Network for Detection of Disorders of Consciousness," in *Chinese Control Conference*, CCC, vol. 2020-July, pp. 7084–7089, IEEE Computer Society, jul 2020.
- [18] Z. S. Zheng, N. Reggente, E. Lutkenhoff, A. M. Owen, M. M. Monti, and Z. Sheng Zheng, "Medical Research Council," 2016.
- [19] P. J. Lang and M. M. Bradley, "Emotion and the motivational brain," jul 2010.
- [20] M. Boly, M. I. Garrido, O. Gosseries, M. A. Bruno, P. Boveroux, C. Schnakers, M. Massimini, V. Litvak, S. Laureys, and K. Friston, "Response to comment on "preserved feedforward but impaired top-down processes in the vegetative state"," dec 2011.
- [21] "fMRI Imaging: How Is an fMRI Done? fMRI Imaging | HowStuffWorks."
- [22] "FMRI Functional Magnetic Resonance Imaging Lab."
- [23] Ç. G. Altıntop, F. Latifoğlu, A. K. Akın, R. İleri, and M. A. Yazar, "Analysis of Consciousness Level Using Galvanic Skin Response during Therapeutic Effect," *Journal of Medical Systems*, vol. 45, jan 2021.

BIBLIOGRAPHY 29

[24] J. Luauté, A. Dubois, L. Heine, C. Guironnet, A. Juliat, V. Gaveau, B. Tillmann, and F. Perrin, "Electrodermal reactivity to emotional stimuli in healthy subjects and patients with disorders of consciousness," *Annals of Physical and Rehabilitation Medicine*, vol. 61, pp. 401–406, nov 2018.

- [25] R. Taherkhani and M. Kia, "Designing a high accuracy 3D auto stereoscopic eye tracking display, using a common LCD monitor," 2012.
- [26] W. K. C. Ting, J. L. Perez Velazquez, and M. D. Cusimano, "Eye movement measurement in diagnostic assessment of: Disorders of consciousness," 2014.
- [27] E. Muller, J. P. Shock, A. Bender, J. Kleeberger, T. Högen, M. Rosenfelder, B. Bah, and A. Lopez-Rolon, "Outcome prediction with serial neuron-specific enolase and machine learning in anoxic-ischaemic disorders of consciousness," *Computers in Biology and Medicine*, vol. 107, pp. 145–152, 2019.
- [28] "HPC Advisory Council Subgroups HPC | GPU."
- [29] D. B. Spalding, "A general purpose computer program for multi-dimensional one- and two-phase flow," *Mathematics and Computers in Simulation*, vol. 23, pp. 267–276, sep 1981.
- [30] A. Beavers, "Alan turing: Mathematical mechanist," Cooper, S. Barry; van Leeuwen, Jan. Alan Turing: His Work and Impact. Waltham: Elsevier, pp. 481–485, 2013.
- [31] S. J. Keegan, *Intelligence in War: Knowledge of the Enemy from Napoleon to Al-Qaeda*. New York: Alfred A. Knopf, 2003.
- [32] A. R. Booker, "Turing and the riemann hypothesis."
- [33] J. P. Bowen, "Alan Turing: Founder of Computer Science," *Lecture Notes in Computer Science (including subseries Lecture Notes in Artificial Intelligence and Lecture Notes in Bioinformatics)*, vol. 10215 LNCS, pp. 1–15, 2016.
- [34] S. J. Russell and P. Norvig, Artificial Intelligence: a modern approach. Pearson, 3 ed., 2009.
- [35] A. L. Samuel, "Some Studies in Machine Learning Using the Game of Checkers,"
- [36] D. Silver, J. Schrittwieser, K. Simonyan, I. Antonoglou, A. Huang, A. Guez, T. Hubert, L. Baker, M. Lai, A. Bolton, Y. Chen, T. Lillicrap, F. Hui, L. Sifre, G. van den Driessche, T. Graepel, and D. Hassabis, "Mastering the game of Go without human knowledge," *Nature* 2017 550:7676, vol. 550, pp. 354–359, oct 2017.
- [37] "Explore the history of machine learning."
- [38] F. Rosenblatt, "THE PERCEPTRON: A PROBABILISTIC MODEL FOR INFORMATION STORAGE AND ORGANIZATION IN THE BRAIN 1," *Psychological Review*, vol. 65, no. 6, pp. 19–27, 1958.

30 BIBLIOGRAPHY

[39] "68747470733a2f2f73656261737469616e72617363686b612e636f6d2f696d616765732f626c6f672f323031352f73 (664×482)."

- [40] "(3) Support Vector Machines: A Visual Explanation with Sample Python Code YouTube."
- [41] A. MUller, Introduction to machine learning with Python: a guide for data scientists. 2017.
- [42] "Taxonomy of machine learning algorithms | Scala for Machine Learning Second Edition."
- [43] B. Sheng, O. M. Moosman, B. Del Pozo-Cruz, J. Del Pozo-Cruz, R. M. Alfonso-Rosa, and Y. Zhang, "A comparison of different machine learning algorithms, types and placements of activity monitors for physical activity classification," *Measurement: Journal of the International Measurement Confederation*, vol. 154, mar 2020.
- [44] "What is a Confusion Matrix? Unite.AI."
- [45] "Confusion matrix Wikipedia."
- [46] D. M. W. Powers, "Evaluation: From Precision, Recall and F-Measure to ROC, Informedness, Markedness & Correlation," *Journal of Machine Learning Technologies*, vol. 2, no. 1, pp. 37–63, 2011.
- [47] H. Gill-Thwaites and R. Munday, "The Sensory Modality Assessment and Rehabilitation Technique (SMART): A Comprehensive and Integrated Assessment and Treatment Protocol for the Vegetative State and Minimally Responsive Patient," http://dx.doi.org/10.1080/096020199389392, vol. 9, pp. 305–320, jul 2010.
- [48] J. Patel, "Unification of machine learning features," in 2020 IEEE 44th Annual Computers, Software, and Applications Conference (COMPSAC), pp. 1201–1205, 2020.