BACHELOR THESIS Artificial Intelligence Radboud University Nijmegen

Diagnosis Systems for Dementia

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Abstract

In 2020 there will be about 42.3 million people worldwide with a form of dementia (Ferri et al., 2006). It seems likely that a fast and precise diagnoses can contribute to better care for the patient and more certainty for the caregiver. Medical diagnosis systems already exist and assist physicians and psychologists worldwide. Still no medical decision support system for diagnosing dementia exists.

The focus of my research will be the design, development and evaluation of decision support system for diagnosing dementia, based on four of the most prominent artificial intelligent techniques. These are:

- Logistic regression
- Nearest neighbor
- Neural networks
- Support vector machines

Because no diagnosis system exists for dementia, the accent of my research will be the exploration of suitable AI techniques for such a system. Furthermore I will look at the practical side of a diagnosis system for dementia by making use of the knowledge of an expert in the field.

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Chapter 1 Introduction

Medical diagnosis systems already exist and assist physicians and psychologists worldwide. Medical diagnosis systems already exist and assist physicians and psychologists worldwide. Still no medical decision support system for diagnosing dementia exists. The focus of my research will be the design, development and evaluation of decision support system for diagnosing dementia, based on four of the most prominent artificial intelligent techniques.

In this introduction the use of decision support systems for medical problems is briefly discussed. The importance of a diagnosis system for dementia will be explained in Section 1.2. Also the epidemiology and diagnosis pipeline of dementia are described. I will discuss more about the psychological diagnosis of dementia in Section 1.1.

1.1 The diagnosis of dementia

1.1.1 Dementia

Dementia is a cluster of different phenomena, particularly the deterioration of memory and other cognitive functions. It is a very frequent condition that strongly increases with age (Jonker, Slaets, & Verhey, 2009).

The term dementia was already used in the Roman era, for patients suffering from 'fever delirium remaining permanent insanity' (Jonker et al., 2009 and Jonker, Verhey, & Slaets, 2001). Dementia was official classified as a syndrome by both Pinel (1818) and Esquirol (1838). Esquirol defined dementia as a psychological decay, caused by a chronic illness of the brain.

In the nineteenth century clinical symptoms and brain pathology became more associated (Jonker et al., 2009). At the time, one believed that arteriosclerosis is the cause of mental deterioration at the end of life. In 1898 Alzheimer had his doubts about this theory and analyzed relative young patients with dementia. He concluded that not all the mental deterioration was caused by vascular changes (Jonker et al., 2009).

Since then al lot of research has been done about the causes of dementia. Nowadays dementia is a general name for a number of symptoms with different causes. According to the latest edition of the Diagnostic and Statistical Manual of Mental Disorders, DSM-IV-TR, the dementia disorders have "the development of multiple cognitive deficits (including memory impairment)" in common (American Psychiatric Association, 2000).

In the DSM-IV-TR dementia section dementia is categorized by cause in six different disorders:

- Dementia of the Alzheimer's Type
- Vascular Dementia
- Dementia Due to Other General Medical Conditions
- Substance-Induced Persisting Dementia
- Dementia Due to Multiple Etiologies
- Dementia NOS (Not Otherwise Specified)

Some of these dementia categories could include more dementia types. For example 'Frontotemporal Dementia', is a dementia type categorized in the DSM-IV-TR under 'Dementia Due to Other General Medical Conditions'.

In the practice of diagnosing dementia Dutch psychologists (e.g. from *de* Zorggroep and UMC St. Radboud) will not diagnose all the types of dementia. Distinguishing between all the types is difficult, because types do not exclude each other. For example a client could have Alzheimer's Disease and Vascular Dementia (Jonker et al., 2009). Moreover for screening clients, the diagnosis of dementia alone could be satisfactory. When needed, a further diagnosis and examination will take place. In dealing with clients who have dementia diagnosing of dementia could be enough.

When a more specific diagnosis is needed, there will be a distinction between four types of dementia. These are the four most common types of dementia (*Dianostiek van dementie*, 2008):

- Alzheimer's Disease ¹
- Vasculair Dementia
- Lewy Body Dementia ²
- Frontotemporal Dementia ²

¹Dementia of the Alzheimer's Type

²The last two, 'Lewy Body dementia' and 'Frontotemporal Dementia', are types categorized in the DSM-IV-TR under 'Dementia Due to Other General Medical Conditions'.

1.1.2 The diagnosis pipeline

"Diagnosis of dementia is a stepwise process that involves examination of patient history and early warning signs, as well as performance screening, assessment of daily functioning, behavioral problems, and caregiver status, with possible referral to specialist clinics for more thorough assessment" (Galvin & Sadowsky, 2012).

In order to come to a diagnosis, the psychologist goes through a diagnosis process. A general procedure is shown in Figure 1.1. The course of the procedure is for example adjusted to the client and his/her surroundings. The order of the different processing stages is variable and some stages are mergeable, replaceable or even removable.

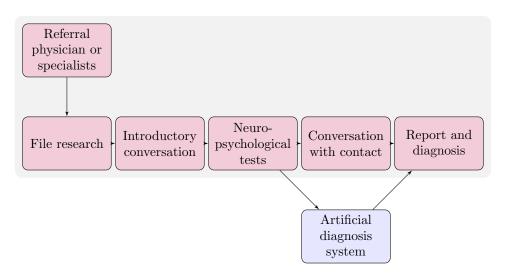


Figure 1.1: General diagnostic procedure and artificial diagnosis system

In the general procedure a client comes to the psychologist after referral. The psychologist begins with a research in the following files of the client: psychological, medical and care file³. This gives background information about the client: there might be physical or mental characteristics, for example a CVA, that are important to take into account in the diagnosis or in dealing with the client.

Next, the psychologist will give a self-introduction and tell more about the course of events. In this conversation the psychologist starts with observing. The psychologist could for example look at the punctually and appearance. Afterward-she neuropsychological tests take place. The psychologist or an assistant will do the

³Two or all files may be combined in a single care file

testing. In the Netherlands (e.g. *De Zorggroep*) the following tests are commonly used: Cambridge Cognitive Examination (CAMCOG) and Mini-Mental State Examination (MMSE). In section 3.1 some of these neuropsychological tests will be described in more detail.

Before making a diagnosis, a contact is consulted. The contact is the spouse, a child or other relative of the client. The contact provides additional information for the psychologist, for example background information about the past of the client. This is also necessary for the hetroanamnesis. When a client is a child, an elderly, a coma patient or mentally disturbed, the client could have a wrong insight in his/her illness and decay. With a hetroanamnesis, the psychologist asks someone else than the client (mostly a close relative) about the illness history of the clients' daily life.

On rare occasions this stage is skipped in the process, but it is not recommended. Most times the information, even if it is limited, helps to make a diagnosis.

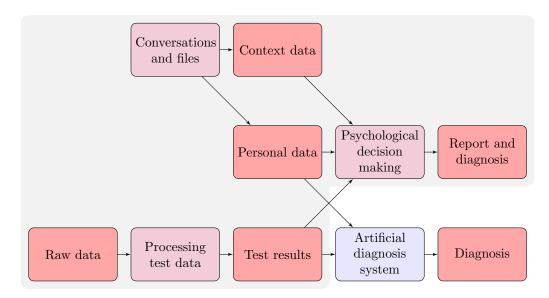


Figure 1.2: Data flow of the human and artificial decision making

At the end the psychologist makes up the report and the diagnosis (for more about the report see section 3.2.2). The conclusion of the report contains the diagnosis and sometimes also recommendations or advice for care staff, specialists or family. In making the diagnosis, the psychologist will look at context, impressions and observations, personal details and the test results. These factors are described in the report. The data flow of the diagnosis process is shown in Figure 1.2.

1.2 Medical decision support systems

In 2020 there will be 42.3 million people worldwide aged more than 60 years, expected with a form of dementia (Ferri et al., 2006). Of those people there will be 6.9 million living in Western Europe. Alzheimer's disease (the most common type of dementia) is one of the most disabling and burdensome health conditions worldwide (Ferri et al., 2006). Most patients with dementia live at home, where family caregiver(s) take care of them (Jonker, Verhey, & Slaets, 2001). However taking care of a demented person contributes to psychiatric morbidity (Schulz & Martire, 2004). It seems likely that a fast and precise diagnoses can contribute to better care for the patient and more certainty for the caregiver.

To come to a better diagnosis process, scientists can analyze, improve and test the human diagnosis process itself with the help of a computer. Many scientists analyzed the human reasoning process in medical diagnosis (Miller, 1994).

But computers can be of more assistance than improving the human diagnosis process. Computers can help the physicians with collecting and processing clinical information (Ledley & Lusted, 1959). Through the years more electronic data is stored and automatically processed. This makes medical decision support systems an addition in improving the accuracy of medical diagnosis (Mangiameli, West, & Rampal, 2004). The prediction is that the use of medical diagnosis systems will tenfold within the decade (Mangiameli et al., 2004).

In Figures 1.1 and 1.2 is the role of the artificial diagnosis system displayed.

Medical diagnosis systems are used to predict/diagnose all kinds of medical problems. Examples are listed in Table 1.1. For more information on the artificial intelligence techniques used for these medical diagnosis systems, see Chapter 2.

CHAPTER 1. INTRODUCTION

	Classifiers	Classifiers
ledical problem	(Mangiameli et al., 2004)	(Kononenko, 2001)
cute myocardial infarction	Logistic Regression,	
	k-Nearest Neighbor,	
	and Multilayer Perceptron	
cute pulmonary embolism	Neural Network	
lood transfusion costs	Neural Network	
oronary artery disease	Logistic Regression	
reast cancer	Logistic Regression,	Symbolic Learning
	k-Nearest Neighbor,	
	Kernel density,	
	Multilayer Perceptron,	
	and other Neural Networks	
olorectal, hepatic and	Multilayer Perceptron	
varian cancer		
ytomegalovirus retinopathy	Multilayer Perceptron	
rug/plasma concentration	Multilayer Perceptron	
allstones	Logistic Regression	
ynecologic cytology smears	Neural Network	
eart diseases of newborn babies		Bayes
iver metastases	Logistic Regression	
ower back disorders	k-Nearest Neighbor,	
	Multilayer Perceptron,	
	other Neural Networks	
ower urinary tract dysfunctions		Symbolic Learning
ymphography		Symbolic Learning
Iortality risk for		
eactive airway disease	Logistic Regression	
epsis	Multilayer Perceptron	
ervere head injury	Kernel density	
pondylarthropathy	Logistic Regression	
urvival of hepatitis		Symbolic Learning
lcers	Logistic Regression	

Table 1.1: Some medical problems use classifiers to predict/diagnose an medical problem

1.3 Research questions

The topic of my research will be the design, development and evaluation of a decision support system for diagnosing dementia based on four of these techniques.

There exists no medical decision support system for diagnosing dementia at present. However a number of promising artificial intelligence techniques may be pursued for the design of such a system. These techniques are shown in Table 1.1 and will be discussed in Chapter 2.

Because no diagnosis system exists for dementia, the focus of my research will be the exploration of suitable AI techniques for a diagnosis system. The techniques I will use for the diagnosis system, are selected for this exploration, although other techniques also might suit for this diagnosis problem.

Furthermore I will look at the practical side of a diagnosis system for dementia, by using knowledge and experience of an expert. When designing the diagnosis system and selecting the attributes, I will keep the practical side of the system in mind.

My research questions are as follows:

- 1. Can artificial intelligence techniques be used for the automated diagnosis of dementia?
 - (a) If so, which techniques are accurate and/or robust?
- 2. How can artificial intelligence techniques be used in practice?
 - (a) Which techniques are consistent with the experts?
 - (b) How much effort does it take to use the artificial diagnosis process in practice?

1.4 Research plan and organization of the thesis

In oder to answer my research questions, I followed the next steps:

- literature study and contact with experts
- data acquisition and pre-processing
- running first round of experiments
- evaluation together with expert
- running second round of experiments
- conclusion and discussion

1.4.1 Preparations for this research

First I reviewed literature of the AI classifiers used in medical diagnosis systems. My conclusions are described in Chapter 2. Furthermore I interviewed experts in the field. These experts have specific knowledge about dementia and the diagnosing process and provided sufficient data for testing.

1.4.2 Data processing

The data that has been made available was submitted by two different health care institutions: *UMC St. Radboud* and *De Zorggroep*.

For combining of the two datasets, I applied relatively straight-forward transformations. Furthermore, I computed attributes, that were not present in one or both datasets, to make the datasets complete. Also attributes were removed from the dataset(s). This were irrelevant attributes that occur in only one of the datasets. In Chapter 3, the data analyzing and pre-processing are described.

After the data was analyzed and preprocessed, I prepared the data for application with the most prominent AI classifiers, in particular a support vector machine, neural network, k-nearest neighbor and logistic regression. I applied the experiments with these classifiers in the state-of-the-art machine learning workbench: Weka. The details of the classifiers and the Weka environment can also be found in chapter 3.

1.4.3 Experimental rounds

The first round of experiments is run to explore is diagnosing problem. After the evaluation of the results of this round, second round is designed to improve the results.

- Round 1 In Round 1, the classifiers were tested in two experimental settings: the cross-validation setting and the split setting. These experimental settings are described in Section 3.6 and the results of the first round are displayed in Chapter 4.
- Round 2A The performances and individual cases of the first round were evaluated with the expert. Round 2A was an experiment round, which anticipated on the evaluation together with the expert. It consisted of testing a dataset with additional attributes.
- **Round 2B** The second round also contained experiment 2B. Round 2B has an attribute reduction of Round 1 with the setting cross-validation.

Round 2, which include the evaluation, the experiments 2A and 2B and their results, are described in Chapter 5.

1.4.4 Conclusion and discusion

In Chapter 6, the conclusions of the experiments and the answers on the research questions will be given.

Chapter 2

The Artificial Intelligence View: Medical Decision Support Systems

In this chapter the most prominent AI classifiers will be discussed. At the end the classifiers used for my research are described.

Machine learning could be the answer for what kind of system makes a good medical diagnosis system for dementia. Machine learning can solve problems in different medical domains with diagnostic and prognostic problems by providing methods, techniques and tools (Magoulas & Prentza, 2001). An advantage for the diagnosis system for dementia is that machine learning technology is suitable for small specialized problems in medical diagnosis (Kononenko, 2001). It is also adequate in cases in which algorithmic solutions are not available or there is a lack of formal models (Magoulas & Prentza, 2001). Kononenko (2001) claims that 'machine learning algorithms were designed and used to analyze medical datasets from the very beginning'.

In his papers (2001, 1998) Kononenko made a comparison of some, as he calls it, "state-of-the-art systems" when applied to a couple of medical diagnostic tasks. The diagnostic tasks included the localization of the primary tumor, the predictions of recurrent breast cancer and the diagnostics of thyroid diseases and rheumatology (Kononenko, Bratko, & Kukar, 1998). An overview is shown in Table 2.1.

Below, the following artificial intelligence techniques are described:

- Logistic regression
- Nearest neighbor
- Neural networks
- Support vector machines

2.1. LOGISTIC REGRESSION

Technique	Classifier	Performance	Transparency	Explanation	Reduction	Missing data handling
Decision tree	Assistant-R	Good	Very Good	Good	Good	Acceptable
	Assistant-I	Good	Very Good	Good	Good	Acceptable
LFC		Good	Good	Good	Good	Acceptable
Bayesian	Naive Bayes	Very Good	Good	Very Good	No	Very Good
classifier	Semi-naive Bayes	Very Good	Good	Very Good	No	Very Good
Neural	Back-	Very Good	Poor	Poor	No	Acceptable
Network	propagation					
k-Nearest Neighbor		Very Good	Poor	Acceptable	No	Acceptable

Table 2.1: The appropriateness of various algorithms for medical diagnosis. (Kononenko, 2011)

2.1 Logistic regression

With the ridge regression theory, standard linear regression can be extended to logistic regression (Cessie & Houwelingen, 1992). Logistic regression predicts the class by using a logistic sigmoid of a linear relationship between the class and the attributes. Logistic regression is a popular technique in biostatistics (Cessie & Houwelingen, 1992).

The logistic model is popular, because the logistic function provides a probability and an S-shape, which gives a description of the combined effect of several attributes (Kleinbaum, Klein, & Pryor, 2002).

The logistic model formula (Kleinbaum et al., 2002):

$$P(y = 1|x_i) = \frac{1}{1 + e^{-\alpha \sum_{j} \beta_j x_{i,j}}}$$

2.2 Nearest neighbor

Nearest neighbor is a machine learning technique that uses the idea that a subject is similar to the objects in the neighborhood of the subject. With k-nearest neighbor, the neighborhood is k objects big.

The k-nearest neighbor algorithm is very simple to implement and often preforms quite good (Russell & Norvig, 2003). Despite that the k-nearest neighbor has a poor knowledge representation, physicians found that the k cases of nearest neighbor were shown as an explanation for the decision (Kononenko, 2001). The best matching case(s), which are the nearest neighbor(s), provide appropriate information for further diagnosis and examination. A disadvantage of nearest neighbor is calculating distances in a large dimension space. The dimension space is the number of attributes. The methods for calculating the distances do not scale well with the dimensions, because the complexity increases rapidly with more attributes (Russell & Norvig, 2003).

The decision rule for nearest neighbor is as follows (Wu, 2012):

 $f(x_i) = \theta(x'),$ $\theta(x_a) = y_a,$ $x' = \arg \min(dist(x_i, x'))$

And in case of k-nearest neighbor:

$$f(x_i) = \theta(x'),$$

$$\theta(x_1, \dots, x_k) = mode(y_1, \dots, y_k),$$

$$x' = \{x' \mid \forall z : dist(x_i, z) \ge dist(x_i, v_k) \ge dist(x_i, v_{k-1}) \ge \dots$$

$$\ge dist(x_i, v_1), v_1 \ne v_2 \ne \dots \ne v_k \ne x_i, x' \in \{v_1, v_2, \dots, v_k\}\}$$

2.3 Neural networks

Artificial neural networks are inspired by neural networks in the brain. A neural network contains input nodes, hidden nodes if necessary and output nodes. Input nodes are used to 'sense' the information, hidden nodes work a sort of extra information gain process and output nodes give the result of the whole process. These nodes are connected by weights, which indicate the strength of the connections between input and output nodes. In case of a multilayer perceptron (MLP) hidden layers, which contain hidden nodes, are added between the input and output layer.

For a large number of medical decision support systems, neural networks have been used, because they have a great predictive power (Mangiameli et al., 2004). Neural networks can map capabilities or associate patterns, generalize, is robust and has high fault tolerance and processes information parallel ant at a high highspeed (Ranjith & Khandelwal, 2011).

Neural networks are not transparent in knowledge representation and explanation of their decisions is not easy (Kononenko, 2001). Symbolic rules are more clear in their decisions. It is possible to extract symbolic rules from a neural network (Kononenko, 2001 and Saito & Nakano, 1988). This technique, symbolic knowledge extraction could be a promising approach to the knowledge acquisition problem (Saito & Nakano, 1988). However the extracted rules tend to be large and complex and offer hardly a useful explanation(Kononenko, 2001).

The decision rule for a two layer neural network (Burges, 1996):

$$f(x_i) = \theta(\sum_{n=1}^{N_s} \alpha_n K(x, s_n) + \beta),$$

$$K = tang(\gamma(x \cdot s_n) + \delta)$$

2.4 Support vector machine

The support vector technique finds the optimal hyperplane separating two classes such that it can be used as a decision boundary (Vuurpijl, Schomaker, & Erp, 2003). The support vector machine is based on the support vector theory of Vapnik.

Support vector machines can handle non-linear solution surfaces using the idea of convolution of the dot-product (Cortes & Vapnik, 1995). Also it is robbust against errors in the training set with the notion of soft margins.

Training of most support vector machines takes a lot of time, especially with a large dimension of space and they are also complex and hard to implement (Platt, 1998). Though by using the sequential minimal optimization algorithm the support vector machine has a better scaling properties, is often faster and easier to implement (Platt, 1998). The support vector machine is an extremely powerful and universal learning machine (Cortes & Vapnik, 1995).

The decision rule for a radial basis function machine (Burges, 1996):

$$f(x_i) = \theta(\sum_{n=1}^{N_s} \alpha_n K(x, s_n) + \beta)$$
$$K = exp(\frac{-||x - s_n||^2}{\sigma^2})$$

2.4.1 Sequential minimal optimization

SMO stands for a support vector machine that uses the sequential minimal optimization algorithm (Platt, 1998). When training a support vector machine, it solves a very large quadratic programming optimization problem. The SMO solves the quadratic programming optimization problem quickly without extra storage (Platt, 1998). Because the SMO scales dimensions very well, the SMO preforms well on large datasets with many attributes (Platt, 1998).

2.5 Techniques used in this research

Nearest neighbor is not difficult to understans and is insightful in how it makes its decisions. A disadvantage of nearest neighbor is the difficulty of calculating useful distances in a large dimension space. In contrast to nearest neighbor, neural networks give no transparent explanation of the decision. Still neural networks are popular, because of their good pattern recognition. The logistic models can provide a useful probability and explain their decisions quite well. An advantage of the support vector machines is that they can handle non-linear solution surfaces. Furthermore the support vector machines is robust against errors in the training set with the notion of soft margins. With the SMO technique, the support vector machines are not slow.

The techniques described in this chapter are prominent and well-known AI techniques for classifying subjects. As shown in Table 1.1, they are also frequently used in solving medical problems. Therefore I decided to apply the following techniques:

- Logistic regression
- Neural network
- Nearest neighbors
- Support vector machine

Chapter 3

Methods

The data acquisition and pre-processing (Section 3.2 and 3.3) will be discussed in this chapter. Because the dataset contains a great part of test results, the neuropsychological tests that occur in my research are discussed (Section 3.1). Section 3.4 is about the first dataset used in this research. This chapter also contains a brief description of the machine learning environment Weka (Section 3.5).

3.1 Neuropsychological tests

The datasets that I used for my research contain attributes (derived) from the test results from neuropsychological tests used by *De Zorggroep* and *UMC St. Radboud* for the diagnosis of, and research on, dementia. The CAMCOG and the MMSE are described in Sections 3.1.1 and 3.1.2. Other neuropsychological tests that occor the *UMC St. Radboud* dataset are listed and described in Section 3.1.3.

3.1.1 Cambridge Cognitive Examination

The CAMCOG is the cognitive section of the Cambridge Mental Disorders of the Elderly Examination (CAMDEX). The CAMDEX is "a comprehensive inventory of information relevant to the diagnoses of dementia" (Lindeboom, Horst, Hooyer, Dinkgreve, & Jonker, 1993). The aim of the CAMDEX is to design a single, standardized instrument for accurate clinical diagnosis of dementia and the dementia classifications: Alzheimer's Disease, Vascular Dementia and Dementia Due to Other General Medical Conditions (Roth, Huppert, Mountjoy, & Tym, 2005).

Compared to other tests the CAMDEX is not as general as the $CARE^4$ and

 $^{^4{\}rm the}$ Comprehensive Assessment and Referral Evaluation

the GMS^5 , but also not as specific as the CERAD⁶ (Lindeboom et al., 1993).

A great advantage of the CAMCOG is that it uses short tasks similar to MMSE (Lindeboom et al., 1993). It takes administering the CAMCOG only 20 to 30 minutes (Koning, Dippel, Kooten, & Koudstaal, 2000). The CAMCOG total score is derived from 60 items from different components (questions and small tasks). The components are distributed over the memory section and the non-memory section. The sections contain two or more scales. Some scales have more subscales. For example the scale 'orientation':

Section	• Scale	\circ Subscale
Memory Section	• Orientation	∘ Time ∘ Place

For the all the scales and subscales see Table 3.1. The distribution of the scores are listed in Appendix A.

Memory Section		Non-memory Section	Non-memory Section	
• Orientation		• Language		
	\circ Time		\circ Comprehension	
	• Place		• Expression	
• Memory		• Concentration/At	tention	
	\circ Remote	• Praxis		
	\circ Recent		\circ Constructie	
	\circ Learning		\circ Ideom/ideat	
		• Calculation		
		• Abstract thinking		
		• Perception		

Table 3.1: Sections, scales and subscales of the CAMCOG

The CAMDEX and CAMCOG can detect dementia at an early stage (Roth et al., 2005). An abnormal total score of the CAMCOG could point out dementia, if other causes are excluded. The valuation of the memory section score is a better indication for (early) dementia of the Alzheimers Type. Also an abnormal score on the non-memory section score could indicate a possible dementia syndrome.

⁵the Geriatric Mental State Schedule

⁶Consortium to Establish a Registry for Alzheimer's Disease

3.1.2 Mini-Mental State Examination

The aim of the Mini-Mental State Examination (MMSE) is the detection of mental status changes, particularly in the elderly and thereby enhances patient care (Cummings, 1993). The MMSE is a structured approach to mental status testing that screens for intellectual impairment (Cummings, 1993).

The MMSE is usually tested together with the CAMCOG. Some components overlap in both tests, and both tests contain the same kind of tasks.

3.1.3 Other neuropsychological tests

A great part of the attributes in the *UMC St. Radboud* dataset describe the physical state of the client. The dataset also contains the results of psychological tests. I will describe the neuropsychological tests of the *UMC St. Radboud* dataset briefly:

Cambridge Cognitive Examination (CAMCOG)

The CAMCOG is a screening tool for dementia (Rijsbergen et al., 2011). See also Section 3.1.1.

Clinical Dementia Rating (CDR)

The CDR is a rating scale for dementia (Rijsbergen et al., 2011).

Fifteen-words test A and B (15-WT)

The 15-WT is a measurement for the short vs. long term memory (Deelman, 1972).

Lawton Instrumental Activities of Daily Living (IADL)

The IADL measures a persons ability to perform tasks in activities of daily life (Graf, 2008).

Mini-Mental State Examination (MMSE)

The MMSE is a measurement for the cognitive grading of patients (Folstein, Folstein, & McHugh, 1975). See also Section 3.1.2.

Neuropsychiatric Inventory (NPI)

The NPI measures the neuropsychiatric symptoms and behavior, that affect most subjects with dementia (Medeiros et al., 2010).

TrailMaking test A and B (TMT)

The TMT measures speed of cognitive processing and executive functioning (Sánchez-Cubillo et al., 2009).

Wechsler Adult Intelligence Scale-III, Digit Span A and B (WAIS-III)

The WAIS contain a series of subtests with different accents on a number of cognitive functions (Sherman & Blatt, 1968). The digit span is one of these subsets. This subset measures primarily the short-term verbal retention (Fenwick & Holmes, 1993).

Only the CAMCOG and MMSE are measures for the determination of dementia. The CDR tests the degree of the dementia and not determination of dementia itself. The 15-WT, IADL, TMT and WAIS-III Digit Span are neuropsychological tests for further or specific insight in the client. The NPI is a test that is applied after the diagnosis of dementia.

3.2 Data acquisition

During the first preparations, I came in contact with two different institutions: care institution *De Zorggroep* and academic hospital *UMC St. Radboud*. Both institutions were interested and assisted by providing data for my research.

3.2.1 UMC St. Radboud data

The University Medical Centre Sint Radboud is an academic hospital related to the Radboud University Nijmegen. In 2012, the UMC St. Radboud had 156.085 clients at the policlinic, 32.163 intramural clients, 49.337 extramural clients, 22.882 elective operations and 24.505 emergency Department visits (*Jaardocument 2011 UMC St. Radboud*, 2011).

The dataset to which I was given access was a part of a dataset which was prepared for (ongoing) research at the UMC St. Radboud and Radboud University. The dataset contains 651 clients with 192 attributes. The list of all the original attributes can be found in Appendix B. A short description of the neuropsychological tests in the UMC St. Radboud dataset was listed in Section 3.1.3.

3.2.2 De Zorggroep data

Stichting Zorggroep Noord- en Midden-Limburg (trade name *De Zorggroep*) is a care institution in Limburg, a province in the Netherlands with several residential and nursing homes and care centra. In 2010 *De Zorggroep* had 2.672 intramural clients, 5.152 extramural clients and 72.137 clients in youth and maternity care (*Jaardocument 2010 Stichting Zorggroep Noord- en Midden-Limburg*, 2010).

In order to create the dataset, I was granted access to anonymous psychological reports of "screeningsonderzoeken". These screening reports have the aim of gaining insight into the cognitive functioning of the client. The psychologist examines the cognitive functions of the client as described in Section 1.1. The reports describe inter alia the details of the neuropsychological test results of the client and the diagnosis. From these reports I could extract the attributes which were important for my research. In Appendix C is an adapted version of a screening report. The general structure of a screening report:

- Personal details such as age and gender
- Examination question
- History of the client
- Heteroanamnese with contact
- Observation
- Diagnostic instrumentation
- Test results
- Conclusion and summary/recommendations

De Zorggroep made data available for research, but it should be noted that these data have been collected for non-research purposes. The screening reports are made for gaining insight into the cognitive functioning of the client for diagnosing dementia. Only the necessary neuropsychological tests to determine the diagnosis will be applied. Furthermore, I only extracted the attributes which were of interest for my research. This results in less attributes, compared to the attributes of UMC St. Radboud dataset (192 attributes). De Zorggroep dataset contains 87 attributes.

Only the CAMCOG and MMSE test results occur both in de *De Zorggroep* and *UMC St. Radboud* data.

For every client I had to read a screening rapport, and extract the attributes. Preparing the reports, for example anonimise, also took time for psychologist. Therefore *De Zorggroep* dataset contains far less subjects compared to *UMC St. Radboud* dataset.

3.3 Pre-processing

For combining the two datasets (*De Zorggroep* and *UMC St. Radboud*), the trivial attributes were made uniform. Furthermore, I applied the following adjustments:

 ${\bf 0}\,$ - No adjustment, both present

Many attributes occur in both data sets. Twelve of these attributes are copied in the combined data set, which forms the basis of the experiments described in this thesis.

```
1 - Transformed
```

Six relatively straight-forward transformations have been performed. These transformations, described in Section 3.3.1, imply:

- categorization of age
- categorization of education

 ${\bf 2}\,$ - Calculated

Twenty of the attributes were not present in one or both datasets and have been computed to make the datasets complete.

- normalization from date-of-birth to age (Section 3.3.1)
- categorization of education (Section 3.3.1)
- CAMCOG scales 'attention' and 'calculation' (Section 3.3.2)
- CAMCOG memory and non-memory section (Section 3.3.2)
- CAMCOG median and threshold of total score and most scales (Section 3.3.3)
- CAMCOG cumulative scores for the memory and non-memory sections (Section 3.3.3)
- 3 Removed

The UMC St. Radboud data set original contained 192 attributes. From *De Zorggroep* screening reports, I extracted 87 attributes. I removed the attributes which were not relevant for my research. This were the attributes that occur in only one of the datasets.

The attributes that still remained athough not present in both datasets, were necessary for the CAMCOG and could be recalculated in the other dataset.

For the visualization of the adjustments I will use flowcharts. These flowcharts will consist of different colors for the *UMC St. Radboud* attributes (blue), *De Zorggroep* attributes (red) and the new or combined attributes (purple). An example is given in Figure 3.1.

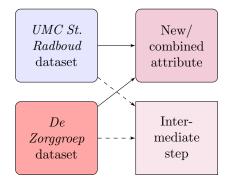


Figure 3.1: Prototype of a flowchart

20

3.3.1 Gender, age and education

In the CAMCOG, the population is distinguished by gender, age category and education level. The CAMCOG uses these characteristics to determine standard scores (Roth et al., 2005). In order to interpret the CAMCOG score, these categories need to be available for computing norms. For these categories, see Table 3.2.

Gender:		Male, Female
Age category:	Ι	$65-69 \text{ years}^7$
	II	70-74 years
	III	75-79 years
	IV	$80-84 \text{ jaar/years}^7$
Dutch	1	Lagere school zes of minder klassen
education level:	2	Voortgezet onderwijs, LO met meer dan 6 leerjaren,
		LBO, ULO, MULO
	3	Middelbaar en hoger onderwijs

Table 3.2: Distinction for norms and scores

The *De Zorggroep* dataset already consisted of an age and age category. But the *UMC St. Radboud* dataset contained none of these attributes. In Figure 3.2 the age and age category attributes are shown. The age and age category were calculated in Excel for all subjects in the *UMC St. Radboud* dataset.

I calculated the age with the following formula:

 $Leeftijd = opname \, datum - geboorte \, datum$

Note that the date of birth is compared to the intake date, instead of the current date. For diagnosing, the age at the examination is important and not the current age.

The age category is calculated according to the CAMDEX- R/N^8 manual (Roth et al., 2005):

$$Age \ category := \begin{cases} 65 - 69 & \text{if} \ age < 70\\ 70 - 74 & \text{if} \ 70 \le age < 75\\ 75 - 79 & \text{if} \ 75 \le age < 80\\ 80 - 84 & \text{else} \end{cases}$$

 $^{^7\}mathrm{The}$ category 65-69 years is for age \leq 69 years, and the category 80-84 years is for age 80 \leq years

 $^{^8 \}rm The \ CAMCOG-R/N$ is the dutch version of the CAMCOG-R

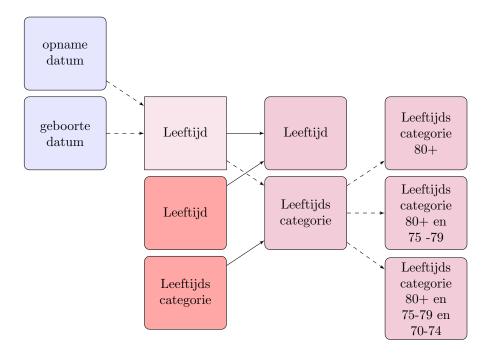


Figure 3.2: Age and Age category attributes evolution

No only the age-related attributes were calculated, the *UMC St. Radboud* dataset misses also the education level attributes. The educational attributes are displayed in Figure 3.3. For the 'education level' attribute, I used the following rule:

$$Education \ level := \begin{cases} 1 & \text{if } education = \text{lagere school zes of minder klassen} \\ 2 & \text{if } ecucation = \text{voortgezet onderwijs, MULO,} \\ & \text{LO met meer dan 6 leerjaren, LBO, ULO} \\ 3 & \text{if } education = \text{middelbaar en hoger onderwijs} \end{cases}$$

The age attributes 'Leeftijds categorie 80+', 'Leeftijds categorie 80+ en 75-79' and 'Leeftijds categorie 80+, 75-79 en 70-74' are added seperatly because the age category is a nominal instead of an ordinal attribute. The same applies to the attribute of the education level, this attribute is split into the two nominal attributes 'Opleidings niveau 1' and 'Opleidings niveau 2'.

3.3.2 CAMCOG test results attributes

Despite that both the datasets contain the test results of the CAMCOG and MMSE, the datasets needed a couple of processing steps. An overview with the evolution of the CAMCOG attributes is shown in Figure 3.4.

3.3. PRE-PROCESSING

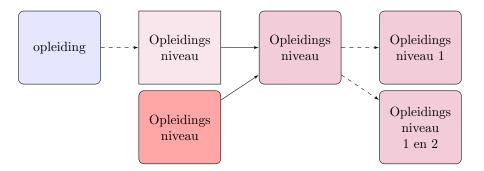


Figure 3.3: Education Level attribute evolution

The Zorggroep dataset contains test results of all the CAMCOG sections, scales and subscales. But the UMC St. Radboud data does not contain the subscales of the CAMCOG. Because the subscale can not be recalculated, these are removed.⁹

Most scales were present in both datasets and needed no processing. But in the $UMC \ St.$ Radboud dataset, the scales 'attention' and 'calculation' are combined into one scale. It is not possible to separate this attribute and I can not recalculate them. Removing would be a waste of valuable data. Therefore the combined scales 'attention' and 'calculation' maintained, and were also combined in *De Zorggroep* dataset.¹⁰

$CAMCOG\,SubA and a cht En Rekenen = A and a cht + Rekenen$

Also the memory section and non-memory section are missing in the UMC St.Radboud dataset. In contrast to the subscales, this section can be calculated. I used the CAMDEX manual (Roth et al., 2005) to calculate the memory section and non-memory section.¹¹

$$\begin{split} CAMCOG\,Geheugensectie &= CAMCOG\,SubGeheugen + CAMCOG\,SubOrientatie\\ CAMCOG\,Niet\,geheugensectie &= CAMCOG\,SubTaal + CAMCOG\,SubPerceptie\\ &+ CAMCOG\,SubPraxis + CAMCOG\,SubAbstractRedenen\\ &+ CAMCOG\,SubAandachtEnRekenen \end{split}$$

 $^{^{9}}$ In Figure 3.4 are these attributes marked with '1'.

 $^{^{10}}$ The combining of scales 'attention' and 'calculation' is shown by number '2' in Figure 3.4

¹¹The calculated memory and non-memory section are marked by number '3' in Figure 3.4

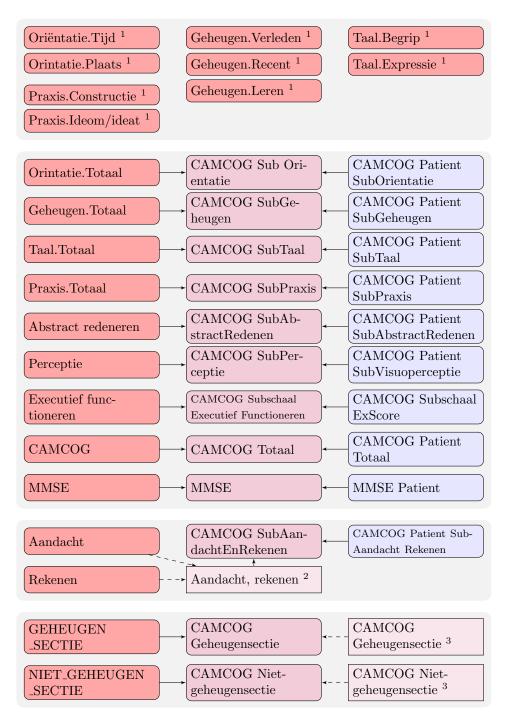


Figure 3.4: The CAMCOG attibutes evolution

3.3.3 CAMCOG norm attributes

The CAMCOG uses a distinction in which the norms are categorized (Roth et al., 2005), see Table 3.2. These are used to calculate individual norms, thresholds and cumulative scores.

The following attributes have a median and a threshold:

- the total score and
- the scales:
 - orientation,
 - language,
 - memory,
 - attention,
 - abstract thinking,
 - and perception.

I calculated both the median and a threshold for the total score and those scales as described in the CAMDEX manual, based on individual characteristics as described above. Even though 'attention' and 'calculation' are combined, I added the norm for 'attention' by itself. For 'calculation' is no norm or threshold (Roth et al., 2005).

The scale 'executive functioning' has no norms for the absence of validation and standards (Roth et al., 2005). Also the scale 'praxis' has no norms calculated (Roth et al., 2005).

The sections memory and non-memory have no median or threshold, but a score. This is a cumulative percentage. The cumulative scores of the memory and non-memory section are also represented in the dataset by the attributes: 'CAM-COG Geheugensectie Score' and 'CAMCOG Niet-geheugensectie Score'. These attributes are calculated according to the tables with the cumulative percentages of the CAMDEX manual (Roth et al., 2005). The table with the cumulative percentages of the memory section in given in Table 3.3.

3.3.4 Dementia types classification

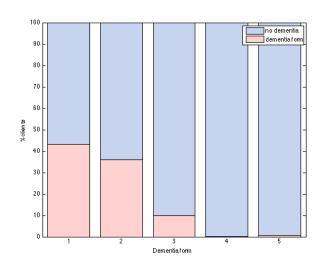
The class 'dementia' was available in both datasets and no further processing was necessary.

Initially the dataset contained also the classes for the four most common types of dementia: Alzheimer's Disease (AD), Vascular Dementia (VS), Lewy Body

Test	Education	Education	Test	Education	Education
score	level 1	levels $2\&3$	score	level 1	levels $2\&3$
19-20	3	1	30	43	24
21 - 22	7	1	31	54	34
23	9	2	32	70	49
24 - 25	13	3	33	85	69
26	15	6	34	94	89
27	18	9	35	99	95
28	23	11	36	100	99
29	32	17	37		100

Table 3.3: The cumulative scores of the memory section determined for each test score and education level

Dementia (LB) and Frontotemporal Dementia (FT). These classes were present in the *UMC St. Radboud* dataset. I extracted more details about the diagnosis from the screening reports. I transformed this information into the four dementia type classes.



(1= Total, 2= Alzheimer's Disease , 3= Vascular Dementia, 4= Lewy Body Dementia, 5= Frontotemporal Dementia)

Figure 3.5: Dementia types balances

As described above, the dataset contained the classes for the four types of dementia. But the balance, the ratio of the subjects of the smallest and largest group, is not sufficient for most of the types. Only the balance of Alzheimers Disease is above 0.3. The balances of the different dementia types are shown in Tabel 3.4 and Figure 3.5.

	TRUE/	Total	UMC St.	De
	total	Dataset	Radboud	Zorggroep
	$\operatorname{subjects}$	balance	balance	balance
AD	241 /670	0.3597	0.3687	0.0526
VS	67 / 670	0.10	0.0968	0.1579
LB	1/670	0.0015	0.0015	0.0
\mathbf{FT}	4 /670	0.0060	0.0061	0.0

Table 3.4: The balances of the dataset for the different dementia types

As mentioned before *De Zorggroep* has giving me access to the screening reports. The determination of the dementia type is not relevant in these reports. In case of positive diagnosis of dementia, further examination could take place. So there could be more cases of AD, VS, LB or FT in the dataset.

The balance of the dementia types is too low and the dementia types could be incomplete. Therefore I decided to skip the classification on the dementia types.

3.4 The dataset for Round 1

The first dataset contains both *De Zorggroep* and the *UMC St. Radboud* dataset. Of the 46 attributes I selected from the screening reports and *UMC St. Radboud* dataset, 38 attributes remained. This attributes are listed in Table 3.5. Some characteristics are shown in Figure 3.6.

		Adjustment
Attribute/class	Scale of measure	number
Geslacht	M,V	0
Leeftijd	numeric	2
'Leeftijdscategorie 80+'	TRUE, FALSE	1
'Leeftijdscategorie $80+$ en $75-79'$	TRUE, FALSE	1
'Leeftijdscategorie $80+$ en 75 - 79 en 70-74'	TRUE, FALSE	1
Leeftijdscategorie	75-79, 80-84, 70-74, 65-69	1
'Opleidingsniveau 1'	TRUE, FALSE	1
'Opleidingsniveau 1 en 2'	TRUE, FALSE	1
Opleidingsniveau	numeric	2
'CAMCOG SubOrientatie'	numeric	0
'CAMCOG SubGeheugen'	numeric	0
'CAMCOG SubTaal'	numeric	0
'CAMCOG SubPraxis'	numeric	0
'CAMCOG SubAandachtEnRekenen'	numeric	2
'CAMCOG SubAbstractRedenen'	numeric	0
'CAMCOG SubPerceptie'	numeric	0
'CAMCOG SubschaalExecutiefFunctioneren'	numeric	0
'CAMCOG Totaal'	numeric	0
'CAMCOG Geheugensectie'	numeric	0
'CAMCOG Niet-geheugensectie'	numeric	2
MMSE	numeric	0
'CAMCOG Totaal Mediaan'	numeric	2
'CAMCOG Totaal Grensscore'	numeric	2
'CAMCOG Geheugensectie Score'	numeric	2
'CAMCOG Niet-geheugensectie Score'	numeric	2
'CAMCOG Orientatie Mediaan'	numeric	2
'CAMCOG Orientatie Grensscore'	numeric	2
'CAMCOG Taal Mediaan'	numeric	2
'CAMCOG Taal Grensscore'	numeric	2
'CAMCOG Geheugen Mediaan'	numeric	2
'CAMCOG Geheugen Grensscore'	numeric	2
'CAMCOG Aandacht Mediaan'	numeric	2
'CAMCOG Aandacht Grensscore'	numeric	2
'CAMCOG AbstractRedenen Mediaan'	numeric	2
'CAMCOG AbstractRedenen Grensscore'	numeric	2
'CAMCOG Perceptie Mediaan'	numeric	2
'CAMCOG Perceptie Grensscore'	numeric	2
Dementie	TRUE, FALSE	0

Adjustment numbers: [0] No adjustment, both present, [1] Transformed, [2] Calculated and [3] Removed

Table 3.5: Attribute list of the first dataset

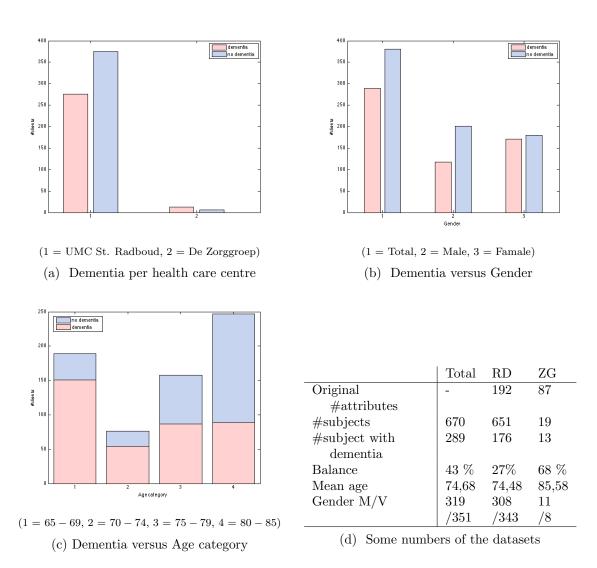


Figure 3.6: Characteristics of the first dataset

3.5 The Weka workbench

For testing classifiers I used version 3.6.7 of the Weka workbench and the Weka manual (Bouckaert et al., 2012). The Weka workbench is a collection of state-of-the-art machine learning algorithms and data pre-processing tools (Frank et al., 2005). Weka supports the whole process of experimental data mining, including preparing the input data, evaluating learning schemes statistically and visualizing both the input data and the result of learning (Frank et al., 2005).



Figure 3.7: Weka logo for version 3.6.7 for Mac

As discussed in Section 2.5, I decided to apply the following techniques: a logistic regression, a neural network, a nearest

neighbors and a support vector machine. Weka has different classifiers for these techniques. The classifiers in Weka, which I applied, are: 'Logistic', 'Multilayer Perceptron', 'NNge' and 'SMO'. The classifiers are described below.

- **Logistic** Classifier for building and using a multinomial logistic regression model with a ridge estimator (Bouckaert et al., 2012).
- Multilayer Perceptron A classifier that uses backpropagation to classify instances (Bouckaert et al., 2012).
- **NNge** Nearest-neighbor-like algorithm using non-nested generalized exemplars (which are hyperrectangles that can be viewed as if-then rules) (Bouckaert et al., 2012).
- **SMO** Implements John Platt's sequential minimal optimization algorithm for training a support vector classifier (Bouckaert et al., 2012).

3.6 Experiments Round 1

In order to answer the research questions, this first round of experiments tests the classifiers on two different characteristics: robustness and comparing rate. The characteristics require a different experimental setting. The settings are described below.

3.6.1 Experimental setting 1, cross-validation

Experimental setting 1 is focused on the first research question:

Can artificial intelligence techniques be used for the automated diagnosis of dementia? If so, which techniques are accurate or robust?

30

In order to test the robustness of the classifiers, I applied ten-fold cross-validation.

Cross-validation is a technique that reduces variability in the performance of a classifier (Russell & Norvig, 2003). The use of cross-validation helps in assessing how the classifier would generalize against an unknown dataset, therefore increasing the reliability of the performance. With k-folds cross-validation, the experiment is run k times. Every time another $(\frac{1}{k})^{\text{th}}$ part is put aside for the test set. The performance of the experiment is with k-folds the average of the k performances.

3.6.2 Experimental setting 2, split

In the second experimental setting I will be able to compare the predictions of the four classifiers, because the test set will be the same for all classifiers. Not only I will be able to compare predictions, the expert can also evaluate prediction of individual subjects. This is necessary for research question 2:

How can artificial intelligence techniques be used in practice? Which techniques are consistent with the experts?

The total dataset was split in a random training set, containing 66% of the subjects, an a test set, containing the other 34% subjects (440 versus 230 subjects). The balance is not held intact precisely, but has just slightly shifted. In the train set is 43% of the subjects is from the dementia class and in the test set this percentage is 42%.

Chapter 4

Results Round 1

4.1 Setting 1

The results of setting 1, the ten-fold cross-validation, are listed in Table 4.1. The parameters of the classifiers used for these performances in my research, are shown in Appendix D. For more details of the results of setting 1, see Appendix E.

Scheme:	MLP ₁	SMO_1	$\operatorname{Logistic}_1$	$k-NN_1$
Correctly classified instances	533 **	541 **	534 **	525 **
	79.55~%	80.75~%	79.70~%	78.36~%
Kappa statistic	0.58	0.61	0.59	0.56
Total number of instances	670	670	670	670

 $^{\star\star}\mathrm{p} < 1\cdot10^{-14}$

Table 4.1: Best results from the Weka classiefiers: MLP, SMO, Logistic and NNge in Round 1 with setting 1: cross-validation

4.2 Setting 2

In this setting, the training and test set are separated. In Table 4.2, the first ten predictions and the performance of the four classifiers are displayed.

The classifiers were *all four* correct in 167 cases (69.58 %), three out of four classifiers were correct in 18 cases (7.50 %), only two correct predictions given by the four classifiers in 6 cases (2.50 %), only one classifier was correct in 11 cases (4.58 %) and in 33 cases (13.75 %) none of the classifiers could find the correct diagnosis. These percentages are displayed in Figure 4.1.

4.2. SETTING 2

	Actual	Predicted	Predicted	Predicted	Predicted	Correct	
Inst	class	MLP_2	SMO_2	$Logistic_1$	$k-NN_2$	predictions	
1	TRUE	TRUE	TRUE	TRUE	TRUE	4	
2	FALSE	TRUE X	FALSE	FALSE	FALSE	3	
3	TRUE	FALSE X	FALSE X	FALSE X	FALSE X	0	
4	TRUE	TRUE	TRUE	FALSE X	FALSE X	2	
5	TRUE	TRUE	TRUE	TRUE	TRUE	4	
6	TRUE	TRUE	TRUE	TRUE	TRUE	4	
7	FALSE	TRUE X	TRUE X	TRUE X	TRUE X	0	
8	TRUE	TRUE	TRUE	TRUE	TRUE	4	
9	TRUE	TRUE	TRUE	TRUE	TRUE	4	
10	FALSE	FALSE	FALSE	FALSE	FALSE	4	
		·			·		
Corre	ectly classified	183 *	182 *	182 *	178 *		
insta	instances 79.57 % 79.13		79.13~%	79.13~%	77.39~%		
Kapp	oa statistic	0.58	0.20	0.57	0.52		
Tota	l number of instances	230	230	230	230		

 $^{\star}\mathrm{p}$ $<1\cdot10^{-9},$ X = incorrect prediction

Table 4.2: Predictions of the first ten instances and performance of the four classifiers in Round 1 with setting 2: split

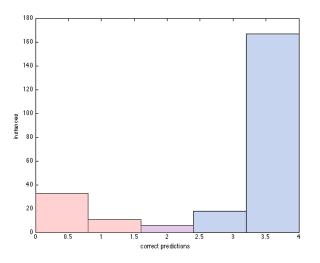


Figure 4.1: Histogram of the correct predictions of the four classifiers

The performance does not improve, when using plurality voting with the four or three best classifiers. See Table 4.3 for the performance, when using plurality voting. An explanation for this could be that in most cases the classifiers make the same prediction, even when they are incorrect (all correct: 68.58 % +all incorrect: 13.75 % = 83.33 %).

MLP ₂	SMO_2	$Logistic_1$	k-NN ₂				
183/230 *	182/230 *	182/230 *	178/230 *				
79.57~%	79.13~%	79.13~%	77.39~%				
	182/230 *						
	79.13~%						
	183/230 *						
79.57~%							

*p <	1	· 1(0^{-9}
------	---	------	----------

Table 4.3: Performance (correct/total instances) of the four classifiers, the three best classifiers combined and all classifiers combined

Chapter 5 Evaluation and Round 2

5.1 Discussion with expert

In the couple of interviews with the expert we discussed several topics, like dementia, the diagnosis process and neuropsychological tests. These were described in previous chapters. The evaluation of the performance and individual subjects was another topics. These and the consequences of incorrect diagnosing will be discussed in the next sections.

5.1.1 Evaluating the performance of the classifiers

The performance of the classifiers is near 80 %. The expert is not surprised to hear this result: the classifier s use only the test results and some personal details (shown in Figure 1.2) and not all the data that the psychologist uses.

The CAMCOG tells something about the cognitive functioning of the client. But low scores are not equal to dementia. As psychologist, you have to exclude other factors and interpret the score to the situation.

Because dementia is deterioration of the cognitive functions, the psychologist is looking for decay. The score of a client is compared to the standard of healthy subjects with the same age, education level, and so on as the clients. Nevertheless the client could have scored, in healthy state, higher or lower than the standard given in the CAMCOG. This means that a high score could be bad, if the client used to be developed above average.

Factors can influence the test results during the examination of the test. A psychologist takes into account whether the client has a handicap. In this way, visual, auditory, phatic or motor impairment issues can make the test more difficult. For example rheumatism can impede writing tasks. Factors like motivation or being tired influence the test results as well.

Some factors influence the CAMCOG specifcally. Depression or even depressive symptoms decrease the CAMCOG score.

There are also other circumstances, like CVA, that decrease the cognitive functions. Then the low scores are not caused by dementia.

The psychologist also look at the illness awareness and illness insight. When a client has illness awareness but no insight, the client knows that he/she has a malfunction. However the client has no insight into the consequences of the illness. For example, a clients, who says:

"I'm forgetful but there is nothing wrong."

is a client with illness awareness but no insight. However when a client says:

"my forgetfulness is a burden for my partner"

tells the clients has both illness awareness and illness insight. It is not possible to have illness insight but no awareness.

5.1.2 Evaluating individual subjects

The instances of the test set of the second setting are both *De Zorggroep* and *UMC St. Radboud* subjects. The expert can get enough details from the original screening reports to reconstruct the diagnosis of *De Zorggroep* subjects. This is not possible with the *UMC St. Radboud* subjects. Of the 230 instances, 9 came from *De Zorggroep*. These nine instances are shown in Table 5.1. The classifiers predicted most instances correct. Only in four cases one or more classifiers predicted incorrect. Instance 3 and 7 were classified incorrectly by all four classifiers. The MLP classifier was wrong with Instance 2. In case of instance 4, the classifiers Logistic and k-NN disagreed with the psychologist.

Inst	Actual class	$\frac{Predicted}{MLP_2}$	Predicted SMO ₂	Predicted Logistic ₁	Predicted k-NN ₂	ID1 in dataset	Sekse	Leeftijds- categorie	Opleidings niveau
1	TRUE	TRUE	TRUE	TRUE	TRUE	1	V	80-84	2
2	FALSE	TRUE	FALSE	FALSE	FALSE	2	V	80-84	2
3	TRUE	FALSE	FALSE	FALSE	FALSE	3	V	75-79	2
4	TRUE	TRUE	TRUE	FALSE	FALSE	4	М	80-84	2
5	TRUE	TRUE	TRUE	TRUE	TRUE	9	V	80-84	1
6	TRUE	TRUE	TRUE	TRUE	TRUE	11	V	80-84	1
7	FALSE	TRUE	TRUE	TRUE	TRUE	14	Μ	80-84	1
8	TRUE	TRUE	TRUE	TRUE	TRUE	15	М	80-84	2
9	TRUE	TRUE	TRUE	TRUE	TRUE	16	V	80-84	2

Table 5.1: The nine instances of *De Zorggroep* in the test set of setting 2

The expert reconstructed the diagnosis of these four instances. The first three were originally diagnosed by a colleague of the expert. Instance 7 was diagnosed by the expert.

Instance 2

Subject 2 was diagnosed with no dementia. The expert confirms this diagnosis, because the cognitive impairment was caused by a CVA.

The majority of the classifiers were indeed correct. The MLP failed because of the CVA. In the Section 5.1.1, the influence of the CVA on the diagnosis was discussed.

Instance 3

The psychologist was not thorough with the examination, for example the expert missed the heteroanamnese in this case. The expert assumed there was not a CVA or other medial deficits and concluded and concluded an early phase of dementia.

Although all classifiers predicted negatively, the diagnosis is still positive. It could be possible that the classifiers did not recognize the dementia, because it is still in an early phase.

Instance 4

Subject 4 was originally diagnosed with dementia. The classifiers did not collective agree: two classifiers agreed and two classifiers disagreed. The experts diagnosed subject 4 with no dementia, but the cognitive disorder NOS^{12} (DSM IV).

The diagnosis was originally incorrect. The classifiers, which predicted initially correct, are now incorrect, and vice versa. This causes the performance of each classifier to change with 0.43%. The performance of the MLP and SMO decreased with 0.43%. The performance of the Logistic an k-NN increased with 0.43%.

Instance 7

The diagnosis of subject 7 is no dementia. The expert confirms this diagnosis. The subject has no dementia, because the cognitive impairment was caused by a CVA. Also the visual impairment (Hemianopsia) and motor impairment probably influenced the test results negatively.

The classifiers probably recognized the cognitive impairment, but were not capable of distinguishing between the CVA and dementia. This is also the case with subject 3 and discussed in Section 5.1.1.

5.1.3 Recommendations

As discussed previously, the CAMCOG is not always conclusive. Human observations could exclude other causes of abnormal cognitive functions. Based on the

¹²Not Otherwise Specified

interview with the expert, I listed the following recommended additional attributes:

- Motivated Is the client motivated during the examination?
- Depressed Has the client depressive symptoms or is the client depressed?
- Sense of failure Is the client aware of failures during the examination?
- Visual impairment Has the client visual impairment?
- Auditory impairment Has the client auditory impairment?
- Phatic impairment Has the client phatic impairment?
- Motor impairment Has the client motor impairment?
- Illness awareness Is the client aware that the he/she is ill?
- Illness insight Has the client insight in the clients' illness?
- CVA Has the client had a cerebrovascular accident (stroke)?

5.1.4 Consequences of incorrect diagnosing

When incorrect diagnosed with dementia, the consequences relate to the cause of the impaired cognitive functions. For example a depression can cause a low cognitive score. When the client is no longer depressed, the cognitive functions return. Client and relatives will still anticipate on dementia, which has negative consequences.

In case of a cognitive disorder, the influence of the diagnosis is not that large. Most cognitive disorders develop dementia and the diagnosis is just pre-mature. Nevertheless the client has no dementia, the cognitive functions are still abnormal. In other situations the client would still be dysfunctional. Worst case scenario is when the client is unnecessary transferred to a psychogeriatric department. Although this situation is not very likely to happen.

The consequences, when incorrectly diagnosed with no dementia, are less dramatic then incorrectly diagnosed with dementia. In case of uncertainty, the psychologist can do a second screening examination, for example a half year later. Because dementia is deterioration of the cognitive functions, the psychologist can compare the two timeframes and look for diminution of cognitive capabilities. The client and relatives will live longer in uncertainty, but are at least not misinformed about the diagnosis.

In conclusion, when it comes to incorrect diagnosing, false positives are less desirable false negatives.

5.1.5 Round 2A: new datasets

The screening reports of *De Zorggroep* contain more information than original extracted attributes. With the help of the expert, the recommended additional attributes were extracted from these reports. Most of the additional attributes

5.1. DISCUSSION WITH EXPERT

could not be found in the UMC St. Radboud dataset and attributes can not be added to the UMC St. Radboud dataset. Round 2A is based only on De Zorggroep dataset, because of the flexibly of this dataset. In Table 5.2, the list of the additional attributes is shown.

TRUE, FALSE
TRUE, FALSE

Table 5.2: Additional attributes for Round 2A

De Zorggroep dataset used in Round 1 contained the 19 subjects. During the research, six other screening reports became available. These reports are not processed in Round 1, but only used in Round $2A^{13}$.

Of the 25 subjects in the new dataset, 17 subjects have dementia. The balance is different from the total dataset used for Round 1 with setting 1 and 2 (17/25 = 0.68 versus 289/670 = 0.43).

For this round, the experimental settings were set according to setting 1 (described in Section 3.6.1). This means I used ten-fold cross-validation. In order to make a more fair compairment possible, not only the dataset with additional attributes was tested. The dataset with the same subjects was tested with the original 38 attributes of Round 1.

5.1.6 Results Round 2A

The results of ten-fold cross-validation with 38 attributes (without additional attributes) are shown in Table 5.3. In Table 5.4, the results with the 48 attributes (with additional attributes) are displayed.

The performance and confusion matrices of the four classifiers on both datasets are combined in Table 5.5.

¹³The reports are also not used in Round 2B

Scheme:	MLP ₃	SMO_3	$\operatorname{Logistic}_1$	k-NN ₃
Correctly classified instances	17 †	18^{+}	20 †	12^{+}
	68~%	72~%	80~%	48~%
Kappa statistic	0	0.38	0.52	-0.24
Total number of instances	25	25	25	25

 $^{\dagger}p > 0.1$

Table 5.3: Best results from the Weka classifiers: MLP, SMO, Logistic and NNge in Round 2A with the original 38 attributes

Scheme:	MLP_4	SMO_3	$\operatorname{Logistic}_2$	$k-NN_4$
Correctly classified instances	17 †	17^{+}	10 †	13^{+}
	68~%	68~%	$40 \ \%$	52~%
Kappa statistic	0	0.26	-0.25	-0.27
Total number of instances	25	25	25	25

 $^\dagger p > 0.1$

Table 5.4: Best results from the Weka classifiers: MLP, SMO, Logistic and NNge in Round 2A with the original and additional attributes

Original 38 attributes

Scheme:	MLP ₃		SMO_3			$\operatorname{Logistic}_1$			k	$k-NN_3$	
	a b	<i>←</i>	a	b	\leftarrow	а	b	←	a	b	→ (
Confusion	17 0	a = True	13	4	a	15	2	a	11	6	a
matrix	8 0	b = False	3	5	b	3	5	b	7	1	ь
		1			I						
	a b	→ (a	b	←	a	b	<i>~</i>	a	b	←
Confusion	17 0	a = True	13	4	a	8	9	a	13	4	a
matrix	8 0	$\mathbf{b} = \mathbf{False}$	4	4	b	6	2	b	8	0	b
Scheme:	MLP ₄		SMO_3		$Logistic_2$		ic_2	k-NN ₄		4	

Original 38 attributes and additional attributes

Table 5.5: Performance and confusion matrices of the four classifiers based on the original 38 attributes dataset and the additional attributes dataset

5.2 Round 2B: attribute reduction

In Round 2B, instead of adding attributes, the performance may improve with a reduction of the attributes. With less attributes, the future space will be smaller.

The attributes with the lowest information gain and gain ratio are best candidates for removal. I used the Weka tools InfoGainAttributeEval, GainRatioAttributeEval and Ranker to calculate the information gain and gain ratio.

InfoGainAttributeEval Evaluates the worth of an attribute by measuring the information gain with respect to the class (Bouckaert et al., 2012).

InfoGain(Class, Attribute) = H(Class) - H(Class|Attribute)

GainRatioAttributeEval Evaluates the worth of an attribute by measuring the gain ratio with respect to the class (Bouckaert et al., 2012).

GainR(Class, Attribute) = (H(Class) - H(Class|Attribute))/H(Attribute)

Ranker Ranks attributes by their individual evaluations. (Bouckaert et al., 2012).

I calculated the *sum* and the *distance* of the info gain and gain ratio to rank the attributes. Top attributes are selected for the new dataset. In Table 5.6 the removed attributes are indicated with a grey colour. The following twenty attributes remain:

- Attribute
- Leeftijd
- Leeftijdscategorie
- Leeftijdscategorie 80+
- Leeftijdscategorie 80+ en 75 -79
- MMSE

- CAMCOG Geheugensectie
- CAMCOG Geheugensectie Score
- CAMCOG Niet-geheugensectie
- CAMCOG Niet-geheugensectie Score
- CAMCOG Perceptie Grensscore
- CAMCOG SubAandachtEnRekenen
- CAMCOG SubAbstractRedenen
- CAMCOG SubGeheugen
- CAMCOG SubOrientatie
- CAMCOG SubPerceptie
- CAMCOG SubPraxis
- CAMCOG SubschaalExecutiefFunctioneren
- CAMCOG SubTaal
- CAMCOG Totaal

Attribute	Info	Gain	Sum	Distance
	gain(1)	ratio (2)	(1) & (2)	(1) & (2)
MMSE	0,35	0,18	0,53	0,40
CAMCOG Totaal	0,32	$0,\!17$	0,50	$0,\!37$
CAMCOG Geneugensectie	0,31	$0,\!18$	$0,\!49$	0,36
CAMCOG SubOrientatie	0,28	$0,\!15$	$0,\!43$	0,32
CAMCOG SubGeheugen	0,28	$0,\!14$	$0,\!42$	0,31
CAMCOG Geheugensectie Score	0,24	$0,\!14$	$0,\!38$	$0,\!28$
CAMCOG Niet-geheugensectie	0,22	$0,\!13$	$0,\!35$	0,26
CAMCOG Niet-geheugensectie Score	0,17	$0,\!11$	0,29	0,21
CAMCOG SubTaal	0,16	0,08	0,24	$0,\!18$
CAMCOG SubschaalExecutiefFunctioneren	0,14	0,09	0,23	$0,\!17$
CAMCOG SubAandachtEnRekenen	0,13	$0,\!10$	0,23	0,16
CAMCOG SubPerceptie	0,14	0,08	0,22	0,16
Leeftijd	0,13	0,07	$0,\!20$	$0,\!15$
CAMCOG SubPraxis	0,12	0,08	$0,\!20$	$0,\!14$
CAMCOG SubAbstractRedenen	0,11	0,07	$0,\!18$	$0,\!13$
Leeftijdscategorie $80+$ en $75-79$	0,08	0,09	$0,\!17$	$0,\!12$
Leeftijdscategorie	0,10	$0,\!05$	0,16	$0,\!12$
CAMCOG Perceptie Grensscore	0,08	0,08	0,16	$0,\!12$
Leeftijdscategorie 80+	0,08	$0,\!08$	$0,\!15$	$0,\!11$
CAMCOG AbstractRedenen Mediaan	0,09	0,06	0,14	0,10
CAMCOG Perceptie Mediaan	0,09	0,06	$0,\!14$	0,10
CAMCOG AbstractRedenen Grensscore	0,09	0,06	$0,\!14$	0,10
CAMCOG Totaal Mediaan	0,07	0,07	$0,\!14$	0,10
CAMCOG Totaal Grensscore	0,07	0,07	$0,\!14$	0,10
Leeftijdscategorie 80+ en 75 - 79 en 70-74	0,07	0,08	$0,\!14$	0,10
CAMCOG Taal Mediaan	0,08	$0,\!05$	$0,\!13$	0,10
CAMCOG Taal Grensscore	0,08	$0,\!05$	$0,\!13$	0,10
CAMCOG Geheugen Mediaan	0,06	0,06	$0,\!13$	0,09
CAMCOG Geheugen Grensscore	0,03	0,06	0,09	0,07
Opleidingsniveau 1	0,03	0,04	0,06	0,05
CAMCOG Aandacht Mediaan	0,01	0,01	0,02	0,02
Opleidingsniveau 1 en 2	0,01	0,01	0,02	0,02
Geslacht	0,01	0,01	0,02	0,01
CAMCOG Aandacht Grensscore	0,01	0,01	0,01	0,01
Opleidingsniveau	0,01	0,01	0,01	0,01
CAMCOG Orientatie Grensscore	0,00	0,00	0,00	0,00
CAMCOG Orientatie Mediaan	0,00	0,00	0,00	0,00

 $(1) \ attribute Selection. In foGain Attribute Eval$

with attribute Selection.Ranker -T $-1.7976931348623157\mathrm{E308}$ -N -1

 $(2)\ attribute Selection. Gain Ratio Attribute Eval$

with attribute Selection.Ranker -T -1.7976931348623157 E308 -N -1

Sum = (1) + (2), Distance =
$$\sqrt{(1)^2 + (2)^2}$$

Table 5.6: The attributes ranked on information gain and gain ratio

5.2.1 Results Round 2B

The results of Round 2B are shown in Table 4.1. For more details see Appendix F.

Scheme:	MLP_5	SMO_4	$\operatorname{Logistic}_3$	$k-NN_5$
Correctly classified instances	533 **	540 **	542 **	530 **
	79.55~%	80.60~%	80.90~%	79.10~%
Kappa statistic	0.58	0.60	0.61	0.57
Total number of instances	670	670	670	670

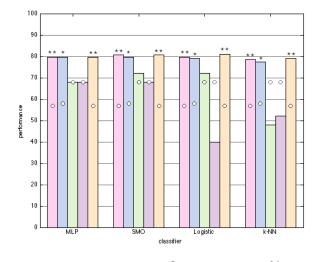
 $^{\star\star}\mathrm{p} < 1\cdot10^{-14}$

Table 5.7: Best results from the Weka classifiers: MLP, SMO, Logistic and NNge in Round 2B

Chapter 6 Conclusion and Discussion

6.1 The results of the AI techniques

In Figure 6.1 the performances of the four classifiers over all rounds are displayed. The results of all rounds are shown in Table 6.1.



 $\circ = \# class / \# total; \ ^{\star}p < 1 \cdot 10^{-9}$, $^{\star\star}p < 1 \cdot 10^{-14}$, else p> 0.1Round 1 setting 1 & 2, Round 2A with & without additional attributes and Round 2B

Figure 6.1: Histogram of the correct predictions of the four classifiers in all rounds

The first results, Round 1 with setting 1, were promising:

MLP: 79.55 %, SMO: 80.75 %, Logistic: 79.70 % and k-NN: 78.36 %.

All classifiers scored significantly better than random at a succes rate of approximately 80% (p < $1 \cdot 10^{-14}$). These results, while using ten-fold cross-validation, were not very different from setting 2 (with 66% split):

6.1. THE RESULTS OF THE AI TECHNIQUES

MLP: +0.03, SMO: -1.62, Logistic: -0.57 and k-NN: -0.97¹⁴.

Those results were also significant ($p < 1 \cdot 10^{-9}$). By using cross-validation when training the classifiers, the variability of the performances is reduced, increasing their robustness. The use of cross-validation helps in assessing how the classifier would generalize against an unknown dataset, so that the performance shows how robust the trained classifier is against unknown datasets.

None of the classifiers outperforms the others. I think the data is not a specific problem solved best by only one classifier technique. Other AI techniques could also be promising as well and should be tested in further research.

The four classifiers do not differ al lot in performance, and give most of the time the same predictions (83.33 %). Only in 12.08 % one classifier disagrees with the others, but in only 7.50 % these are correct.

The performance does not improve, when using plurality voting with the four or three best classifiers. An explanation for this could be that in most cases the classifiers make the same prediction, even when they are incorrect.

In the evaluation together with the expert, the performance and predictions of the classifiers were discussed. The expert stated that the input missed the observations and data to interpret the CAMCOG scores. For example all classifiers could predict dementia, but the client could have had a CVA. The CVA would cause low cognitive functions and therefore the client has no dementia.

It is likely that the classifiers learned to identify the low cognitive functions, but failed to learn all the causes of the CAMCOG score. This would explain way all four classifiers fail in predicting the same subjects. By adding the recommended attributes to the dataset, the performance should increase. The new dataset was small (25 subjects), and probably too small for training the classifiers. In this Round, 2A, the results are scattered and insignificant.

In order to improve the performance in another way than adding attributes, Round 2B was designed to remove attributes. These results were significant and generally the same at the results of Round 1. The differences are slightly better:

MLP: +0.0, SMO: -0.15, Logistic: +0.40 and k-NN: +0.74 ¹⁵.

The performance could be influenced by the variation in psychologists, which diagnosed the patients. But I think this influence could be insignificant. The subjects in the *UMC St. Radboud* dataset are multiple times diagnosed with dementia. Only *De Zorggroep* screening reports are made by one psychologist. This is 2.84 % of the total dataset.

 $^{^{14}}$ difference is 'Round 1, setting 1 - Round 1, setting 2',

 $^{^{15}}$ difference is 'Round 1, setting 1 - Round 2B',

Round 1, setting 1	MLP_1	SMO_1	$\operatorname{Logistic}_1$	$k-NN_1$
Correctly classified instances	533 **	541 **	534 **	525 **
	79.55~%	80.75~%	79.70~%	78.36~%
Kappa statistic	0.58	0.61	0.59	0.56
Total number of instances	670	670	670	670
	1			
Round 1, setting 2	MLP_2	SMO_2	$\operatorname{Logistic}_1$	$k-NN_2$
Correctly classified instances	183 *	182 *	182 *	178 *
	79.57~%	79.13~%	79.13~%	77.39~%
Kappa statistic	0.58	0.20	0.57	0.52
Total number of instances	230	230	230	230
Round 2A,				
without additional attributes	MLP ₃	SMO_3	$Logistic_1$	k-NN ₃
Correctly classified instances	17 †	18 [†]	20 †	12^{+}
	68~%	72~%	80~%	48~%
Kappa statistic	0	0.38	0.52	-0.24
Total number of instances	25	25	25	25
Round 2A,				
with additional attributes	MLP ₄	SMO_3	$Logistic_2$	$k-NN_4$
Correctly classified instances	17 †	17 †	10 †	13 [†]
, i i i i i i i i i i i i i i i i i i i	68~%	68~%	$40 \ \%$	52~%
Kappa statistic	0	0.26	-0.25	-0.27
Total number of instances	25	25	25	25
Round 2B	MLP_5	SMO_4	$\operatorname{Logistic}_3$	$k-NN_5$
Correctly classified instances	533 **	540 **	542 **	530 **
	79.55~%	80.60~%	80.90~%	79.10~%
Kappa statistic	0.58	0.60	0.61	0.57
Total number of instances	670	670	670	670
	1			

$$\label{eq:prod} \begin{split} ^{\dagger}p &> 0.1 \\ ^{\star}p &< 1 \cdot 10^{-9} \\ ^{\star\star}p &< 1 \cdot 10^{-14} \end{split}$$

Table 6.1: Results of the four classifiers in all rounds

6.2 The artificial diagnosis system in practice

The AI techniques for a diagnosis system for dementia used in this research were designed to use available data from two different institutions: care institution *De Zorggroep* and academic hospital *UMC St. Radboud.* In this way, the system would fit in the human diagnosis process pipeline and psychologists would not need to examine more tests than usual.

The attributes used in this research come from standard data in the human psychological diagnosis process. Therefore using a system based on these attributes would not require more effort then filling in a form (like a screening report). When the processing of the test data is automated, a diagnosis system could fit even better¹⁶. Only the observed data would need to be added, just by clicking some checkboxes.

In this research I skipped the classification of the four most common dementia forms. The ratio of dementia form versus non-dementia was too low and *De Zorggroep* dataset could be incomplete. But more important, in practice is the diagnosis in dementia form preserved for further examination. When a further diagnosis is necessary, another specialist system could examine the client in case of dementia.

It is possible to classify Alzheimer's Disease on solely the UMC St. Radboud dataset. Alzheimers Disease is present in 36.87% of the subjects. The other forms have balances lower than 0.10: Vascular Dementia (0.0968), Lewy Body Dementia (0.0015) and Frontotemporal Dementia (0.0061). I recommend you should acquire a new dataset, when looking at a new diagnosis system specialized in dementia forms.

Especially the screening reports of *De Zorggroep* contain valuable information for a diagnosis system for dementia, because they consist of all the relevant information on which the psychologist makes a diagnosis. The *UMC St. Radboud* contains a lot of other information used for academic research. The disadvantage of *De Zorggroep* dataset is that it takes a lot of work to extract the attributes, and the dataset contains only 25 subjects, which is actually too small for testing representatiely.

Therefor I recommend to use a larger dataset based on the screening reports for further research into a diagnosis system for dementia.

When using a diagnosis system in practice, 80% is not enough. One incorrect

 $^{^{16}\}mathrm{For}$ the 'processing of the test data' and 'artificial diagnosis system' process steps look at Figure 1.2

diagnosis on every five cases would increase the trust in the diagnosis system. But I believe that with further research in possible AI techniques and with a bigger dataset of screening reports a diagnosis system could be trained that would assist and improve the diagnosis of dementia.

6.2.1 Final considerations

I am very grateful for the two institutes that have provided the data for my research. Since obtaining real patient data requires the full cooperation of the medical responsible personnel and careful attention for ethical procedures, the data acquisition and pre-processing process should not be underestimated. However, both the *UMC St. Radboud* and *De Zorggroep* were very interested in the outcomes of this research and therefore were able to provide me with this relevant data set.

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First of all, I would like to give my special thanks to the expert, Josma Kunneman, who showed me the real world. A world where AI could contribute to. She, as an expert, gave me the possibility to get some insight in the psychological and practical side of my field. I am grateful for the work and effort she made to provide me relevant literature, contacts, screening reports, interviews and other information. Furthermore she inspired me to research this thesis topic.

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I also want to thank Martijn Lappenschaar for providing the *UMC St. Radboud* dataset, as well as Marcel van Gerven for commenting on the draft version and being the second reader of this thesis.

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Appendix A

In this appendix, the sections of CAMCOG are displayed with their maximum score.

Section	Max score
CAMCOG Total score	107
MMSE	30
Executive functioning	28

Memory section			37
· ·	Orientation		10
		Time	5
		Place	5
	Memory		27
		Remote	6
		Recent	4
		Learning	17
Non-memory sec	70		
i	Language		30
		Comprehension	9
		Expression	21
	Attention/Concen	tration	4
	Praxis		12
		Construction	6
		Ideom/ideat	6
	Calculation	· · ·	5
	Abstract thinking		8
	Perception		11

Appendix B

This is complete list of all the original attributes of the UMC St. Radboud dataset (original file: Geheugenpoli.xls):

ID1

Catogrie geslacht1 gewicht SamenstellingHuishouden GeboorteLandPatient Woning opleidingAnders HoogstebetrekkingPatient Thuiszorg PGDagopvang Voorgescheidenis LichtOZ Dementie Diagnose Patient MDO1AD DLB Dementie DementieDiagnoseMDO2Akkoord DD2DD4 DD1PsychiatrischeAndere DD2DementieAnders DD2NeurologischeAndere DD3PsychiatrischeAndere CDRScoreMDO1 SenielPreseniel SenielPreseniel8 AnderePsychaitrischeDiagnose AnderDementieDiagnoseMDO2 AndereNeurologischDiagnoseMDO2 Hypertensie HypertensieSindsMaand Diabets ÷

opname datum geboortedatum lengte woonsituatie SamenstellingHuishoudenAnders burgerlijke staat opleiding aantal jaar onderwijs Hoogstebetrekkingpartner Dagopvang SomDagbehandeling Anamnese Medicatie Dementie Diagnose Patient MDO2VaD FTD DementieDiagnoseMDO1Akkoord DefintieveDiagnose DD3DD1DementieAnders DD1NeurologischeAndere DD2PsychiatrischeAndere DD3DementieAnders DD3NeurologischeAndere CDRScoreMDO2 SenielPreseniel4 AnderDementieDiagnose AndereNeurologischDiagnose AnderePsychaitrischeDiagnoseMDO2 WanneerZijnDeGeheugenProblemenBegonnen HypertensieSindsDag HypertensieSindsJaar **DiabetesSindsDag**

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DiabetesSindsMaand CardiovasulaireZiekten RokenSinds **RokenSindsMaand** RokenStop RokenStopMaand RokenHoeveel RokenHoeveelperTijd HuidigeAlcoholinname AlcoholHoeveelheidPerTijd DepressieDag DepressieJaar Overgewicht HypercholHypertriDag HypercholHypertriJaar Vascular Respiratoir BovensteTractusDigestivus Lever UrogenitaleSysteem Neurologisch PsychiatrischeZiekten aantal catogorie aantal cat in 4 Hart 4 **MMSEPatient** GDS15Patient **CAMCOGPatientTotaal CAMCOGPatientSubOrientatie CAMCOGPatientSubTaal** CAMCOGPatientSubGeheugen CAMCOGPatientSubAandachtRekenen**CAMCOGPatientSubPraxis** CAMCOGPatientSubAbstractRedenen**CAMCOGPatientSubVisuoperceptie** CAMCOGSubschaalExScore 15WoordVersie 15WoordReeks2 15WoordReeks4 15 WoordReeksTScore15 WoordRecallTscorePatient15WoordRecognitionPatientAfgenomen TrailMakingAPatient TrailMakingBPatient CijferreeksenAPatient SchaalscoreCijferreeksen

DiabetesSindsJaar Roken RokenSindsDag RokenSindsJaar RokenStopDag RokenStopJaar RokenEenheid Alcohol AlcoholHoeveelheid Depressie DepressieMaand FamiliareBelasting HypercholHypertri HypercholHypertriMaand Hart Haematopoetisch OgenOrenNeusPharynxEnLarynx **OndersteTractusDigestivus** Nieren NeuromusculiareSysteem Endocrine EnMetabole System Mammaetotaal score aantal cat in 3 Hart 3 poli MMSEPatientScore GDS15PatientScore **CAMCOGPatientMaxScore CAMCOGPatientSubOrientatieMax CAMCOGPatientSubTaalMax** CAMCOGPatientSubGeheugenMax CAMCOGPatientSubAandachtRekenenMaxCAMCOGPatientSubPraxisMaxCAMCOGPatientSubAbstractRedenenMax**CAMCOGPatientSubVisuoperceptieMax** CAMCOGSubschaalExMaxScore 15WoordReeks1 15WoordReeks3 15WoordReeks5 15 WoordRecallPAtient15WoordRecognitionPatient 15woordPatientAfgenomenVersie **TscoreTrailMakingAPatient TscoreTrailMakingBpatient** CijferreeksenBPatient TinettiBalansPatient

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56

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TinettiBalans Patient Max Score ${\it TinettiLopenPatientMaxScore}$ BarthelPatiënt LawtonPatient BMIPatient RRDiastolischPatient NaasteMantelzorger $\label{eq:Anderemantel} Anderemantel zorger DAN aaste Wie$ GeslachteNaaste RelatiePatientNaasteMannelijk ZBINaaste NPI NPIEmo AanvullendeDiagonostiek KlachtenWanneerMaand PatientRoute MTARechts

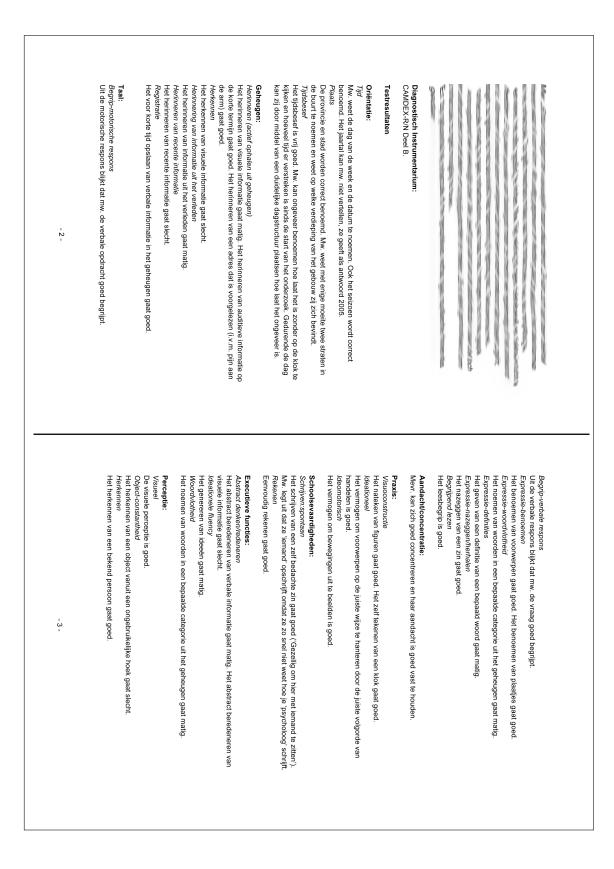
:

TinettiLopenPatient TimedUpAndGoPatient BarthelPatiëntMaxScore LawtonPatientMaxScore **RRSystolischPatient** NaasteInBeeld $\label{eq:andereMantelzorgerDanNaaste} AndereMantelzorgerDanNaaste$ GeboortedatumNaaste RelatiePatientNaasteVrouwelijk RelatiePatientNaasteAndersZBINaasteMaxScore NPIMaxScore NPIEmoMax KlachtenWanneerDag KlachtenWanneerJaar MTALinks TscoreTrailMakBtovApat

Appendix C

An example of a screening rapport of *De Zorggroep*. First, the rapport was made anonymous by the psychologist. Personal information about the psychologist is removed due to privacy reasons.

Mathematical and the second and the	De Zuggroep is de handelsneam van Stichting Zuggroep Noord- en Midden-Limburg
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Condercies Ammunication Maximumication Maximumication Condentiant	SUDWA-	Overzicht testresuitaten				
Tidnifatie Tigl (139 tm 143) 5 4 5/3 5/3 Team (14) Team (14) 5 5 4 5/3 5/3 Team (172 tim 15) Team (171) 6 3 5/3 7/3/10 Scheugen Verheden (166 tim 171) 6 3 3/3 3/3 Recent (172 tim 15) 10 10 10 10 10/3 Team (164, 165, 166) 17 10 13/10 * * Team (164, 165, 165, 166) 17 10 13/10 * * Team (164, 165, 165, 165, 165, 165, 165, 165, 165	Onderdeel		Maximum- score		Mediaan/ grens	Onder
Flaats (144 tim 146) 5 5/3 6/3 7 Ösheugen Fotaal 1008 3 103 10 Totaal Resert (172 tim 173) 1	Oriëntatie	Tijd (139 t/m 143)	5	4	5/4	0
		Plaats (144 t/m 148)	5	5	5/3	
$ \begin{array}{ $		Totaal	10	6	10/8	
$ \frac{\text{Recent (172 (m175)}}{1 \text{ clail}} = \frac{4}{217} = \frac{3}{212} = \frac{43}{2170} = \frac{3}{217} = \frac{13}{10} = \frac{13}{10}$	Geheugen	Verleden (166 t/m 171)	9	e	5/3	
$ \frac{ \text{Leffent (164,165,164)}}{ \text{Leffent (164,165,164)}} \frac{17}{27} \frac{10}{23} \frac{13,10}{23} \frac{1}{23} \frac$		Recent (172 t/m175)	4	e	4/3	
Iotaal Iotaal 27 16 2320 ** aleugensectia 37 25 perc. ** * * *		Leren (164,165,196)	17	10	13/10	
alterugensectie 37 25 perc. Tail Engrip (149 Um 155, 161, 162) 9 8 91 98 91 98 91 98 91 98 91 98 91 98 91 98 91 98 91 98 91 98 93 9		Totaal	27	16	23/20	*
Begrip (149 tm 155, 181, 182) 9 8 9/8 * Total Expressio (165 tm 165) 21 19 272 2725 Total (178, 179) 7 6 6/3 5 5 Praxis Constructile (133, tm 169) 6 5 6/4 5 6/4 5 6/4 5 6/4 5	Seheugensectie		37	25	perc.	
Expressie (156 tim 163) 21 19 1 Totaal 30 27 27/25 27/25 Paxis Constructile (183 tim 166) 6 6/3 6 6/3 Paxis Constructile (183 tim 166) 6 6 6/3 6 6/3 6 6/3 6 6/3 6 6/3 6 6/3 6 6/3<	Taal	Begrip (149 t/m 155, 181, 182)	6	80	9/8	*
Total 30 27 27/25 Aundacht (178,179) 6 6/3 6 6/3 Paxis Ideom/deat (183,191/m 183) 6 5 6/		Expressie (156 t/m 163)	21	19		
andacht (178, 179) 7 6 6/3 6 Praxis Constructie (133, 11m 133) 6 5 6 6/4 7 Pastis Ideomidati (133, 11m 133) 6 5 5 6 6/4 7 Retenen (194, 195) 12 12 14 6 7 4 Persopia (194, 195) 139 2 2 2 7 4 Persopia (194, 195) 139 200) 8 3 5/3 7 4 Persopia (194, 195, 199, 200, 200a, 200b) 28 16 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 4 7 4 7 4 7 4 7 4 7 4 7 4 7 4 7 4 7 4 7 4 7 4 7 4 7 7		Totaal	30	27	27/25	
Taxis Constructie (183 tim 186) 6 64 64 Texis Ideom/deat (183 tim 133) 6 5 6/5 6/5 Ideom/deat (183 tit im 133) 6 7 12 7 2 <	Aandacht	(178,179)	7	9	6/3	
Ideom/Ideat (198, 191 tm 133) 6 5 6/5 <td>raxis</td> <td>Constructie (183 t/m 186)</td> <td>9</td> <td>9</td> <td>6/4</td> <td></td>	raxis	Constructie (183 t/m 186)	9	9	6/4	
Total 12 11 1 Rekenen (194, 195) 2		Ideom/ideat (189.191 t/m 193)	9	2	6/5	
cekenen (194, 195) 2 2 7/4 ubstract redemenen (197, tm 200) 8 6 7/4 5/3 ubstract redemenen (197, tm 200) 8 3 5/3 5/3 5/3 uerceptie (201, 202) 8 16 5/3 perc. perc. perc. perc. perc. perc. perc.		Totaal	12	11		
Obsiziant leadeneren (187 tm. 200) 6 7/4 7/4 Verceptie (201, 202) 8 3 5/3 5/3 Verceptie (201, 202) 8 3 5/3 5/3 5/3 Verceptie (301, 202) 188, 197, 198, 198, 198, 198, 199, 200, 208 67 55 perc. Recursifie (158, 197, 198, 198, 200, 208, 208, 208) 28 16 776 Institutioner (104, 48, 68, 176, 179, 10m) 30 26 7776 AMSC (139, 118, 157, 179, 10m) 30 26 7776 AMSC (134, 118, 157, 179, 10m) 30 26 7776 AMSC (134, 118, 157, 179, 10m) 30 26 7776 Institutioner 30 26 776 2776 Institutioner 30 26 101, 175 279	Rekenen	(194, 195)	2	2		
errore for an international and an international and an international and an international and and an international and	hetract redenare		ι α	ı «	VIL	
Intercent Normalization F 5 Perc. Perc. ehengensectie (158, 197, 198, 198, 200, 200a, 200b) 28 16 Perc.	arcentia		α	» «	512	
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xecutef (158.197.198.198.200.200a.200b) 28 16 10 Amedioneen 104 80 91/78 5 Amedioneen 30 26 91/78 5 IMSE (139 tum48, 156.179 tum 30 26 91/78 5 Immorrating endormating endormating enconcludeerd worden dater sprake is van een beginnendementing endormerities beeld.	liet- eheugensectie		67	55	perc.	
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AMCOG 104 80 91/78 MSE (139 km/48, 156, 163, 176, 179 km) 30 26 7 Ist, 133, 157 en 189) 30 26 7 7 normgroep: vrouw left)(jdf, 75-79 op1.nix, 2 2 4 amerivating en conclusie: op1.nix, 2 0 1 1 amerivating en conclusie: en beginnend 4 4 4 ennolieel beeld. 	Inctioneren	10011011011001000000000	07	2		
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normgroep: vrouw leeflijd: 75-79 opl.niv: 2 amerivatting en conclusie: it he huidige onderzoek kan geconcludeerd worden dat er sprake is van een beginnend amentieel beeld. et vriendelijke groet.	IMSE	(139 t/m148, 156, 163, 176, 179 t/m 181, 183, 187 en 188)	30	26		
amerivating en conclusie: amerivating en conclusie: it het huidige onderzoek kan geconcludeerd worden dat er sprake is van een beginnend amentieel beeld. et vriendelijke groet,			C			
thet huidige onderzoek kan geconcludeerd worden dat er sprake is van een beginnend ementieel beeld. et vriendelijke groet,	amenvatting en	reciulu: 73-73 conclusie:	opi.illv. z	_		
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å	ementieel peeld.					
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4	let vriendelijke gro	let,				
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Appendix D

The four classifiers are identified a index. In this appendix, the parameters of these classifiers are described.

$\begin{array}{c} \mathbf{MLP}_1\\ \mathbf{MLP}_2\\ \mathbf{MLP}_3\\ \mathbf{MLP}_4\\ \mathbf{MLP}_5 \end{array}$	functions. Multilayer Perceptron -L 0.0050 -M 0.2 -N 750 -V 0 -S 0 -E 20 -H 2 functions. Multilayer
\mathbf{SMO}_1	functions.SMO -C 1.0 -L 0.0010 -P 1.0E-14 -N 0 -V -1 1 -K
\mathbf{SMO}_2	functions.supportVector.PolyKernel -C 250007 -E 1.0 functions.SMO -C 1.0 -L 0.0010 -P 1.0E-16 -N 0 -V -1 1 -K functions.supportVector.PolyKernel -C 250007 -E 1.1
\mathbf{SMO}_3	functions. SMO -C 1.0 -L 0.0010 -P 1.0E-12 -N 2 -V -1 1 -K
\mathbf{SMO}_4	functions. supportVector. PolyKernel -C 250007 -E 1.2 functions. SMO -C 1.0 -L 0.0010 -P 1.0E-20 -N 0 -V -1 1 -K functions. supportVector. PolyKernel -C 250007 -E 1.3
$\mathbf{Logistic}_1$	functions.Logistic -R 1.0E-8 -M -1
$\operatorname{Logistic}_2$	functions.Logistic -R 1.0E-10 -M -2
$\mathbf{Logistic}_3$	functions.Logistic -R 1.0E-6 -M -3
$\begin{array}{c} \mathbf{Nnge}_1\\ \mathbf{Nnge}_2\\ \mathbf{Nnge}_3\\ \mathbf{Nnge}_4\\ \mathbf{Nnge}_5 \end{array}$	rules.NNge -G 20 -I 10 rules.NNge -G 24 -I 6 rules.NNge -G 2 -I 1 rules.NNge -G 5 -I 1 rules.NNge -G 10 -I 3

Appendix E

This appendix contains the results of Round 1, setting 1. The full dataset with 38 attributes is classified with ten-folds cross-validation.

MLP: Multilayer Perceptron

Summary				Cor	nfusion Matrix
Correctly Classified Instances Incorrectly Classified Instances Kappa statistic Mean absolute error Root mean squared error Relative absolute error Root relative squared error Total Number of Instances	$533 \\ 137 \\ 0.58 \\ 0.26 \\ 0.38 \\ 53.41 \% \\ 77.27 \% \\ 670$	$\begin{array}{c} 79.55 \ \% \\ 20.45 \ \% \end{array}$	a 221 69	b 68 312	$\leftarrow classified as$ $a = TRUE$ $b = FALSE$

Detailed Accuracy By Class

	TP Rate	FP Rate	Precision	Recall	F-Measure	ROC	Class
						Area	
	0.765	0.181	0.762	0.765	0.763	0.872	TRUE
	0.819	0.235	0.821	0.819	0.82	0.872	FALSE
Weighted Avg.	0.796	0.212	0.796	0.796	0.796	0.872	

SMO: Suport Vector Machine

Summary

Correctly Classified Instances	541	80.75~%	
Incorrectly Classified Instances	129	19.25~%	
Kappa statistic	0.61		
Mean absolute error	0.19		-2
Root mean squared error	0.44		_ <u>ک</u>
Relative absolute error	39.24~%		į
Root relative squared error	88.60~%		
Total Number of Instances	670		

Confusion Matrix

a	b	\leftarrow classified as
218	71	a = TRUE
58	323	$\begin{aligned} \mathbf{a} &= \mathbf{TRUE} \\ \mathbf{b} &= \mathbf{FALSE} \end{aligned}$

62

	TP Rate	FP Rate	Precision	Recall	F-Measure	ROC	Class
						Area	
	0.754	0.152	0.79	0.754	0.772	0.801	TRUE
	0.848	0.246	0.82	0.848	0.834	0.801	FALSE
Weighted Avg.	0.807	0.205	0.807	0.807	0.807	0.801	

Logistic: Logistic Regression

Summary

534	79.70~%			
136	20.30~%			
0.59			h	\leftarrow classified a
0.25			-	
0.20		219	70	a = TRUE
0.0.		66	315	b = FALSE
51.88~%		00	010	0 - IIIDD
74.22~%				
670				
	$\begin{array}{c} 136\\ 0.59\\ 0.25\\ 0.37\\ 51.88\ \%\\ 74.22\ \%\end{array}$	$\begin{array}{cccc} 136 & 20.30 \ \% \\ 0.59 & \\ 0.25 & \\ 0.37 & \\ 51.88 \ \% & \\ 74.22 \ \% & \end{array}$	$\begin{array}{cccccc} 136 & 20.30 \ \% \\ 0.59 & & \\ 0.25 & & \\ 0.37 & & \\ 51.88 \ \% & & \\ 74.22 \ \% & & \\ \end{array}$	$\begin{array}{cccc} 136 & 20.30 \ \% \\ 0.59 & & \\ 0.25 & & \\ 0.37 & & \\ 51.88 \ \% & & \\ 74.22 \ \% & & \\ \end{array} \qquad \qquad$

Detailed Accuracy By Class

	TP Rate	FP Rate	Precision	Recall	F-Measure	ROC	Class
						Area	
	0.758	0.173	0.768	0.758	0.763	0.884	TRUE
	0.827	0.242	0.818	0.827	0.822	0.884	FALSE
Weighted Avg.	0.797	0.212	0.797	0.797	0.797	0.884	

k-NN: Nearest Neighbor

Summary

Correctly Classified Instances Incorrectly Classified Instances	$525 \\ 145$	$\begin{array}{c} 78.36 \ \% \\ 21.64 \ \% \end{array}$			
Kappa statistic Mean absolute error	$0.56 \\ 0.22$		a	b	\leftarrow classified as
Root mean squared error	$0.22 \\ 0.47$		208	81	a = TRUE
Relative absolute error	44.11 %		64	317	$\mathbf{b} = \mathbf{FALSE}$
Root relative squared error Total Number of Instances	$9.93\ \%\ 670$				

Detailed Accuracy By Class

	TP Rate	FP Rate	Precision	Recall	F-Measure	ROC	Class
						Area	
	0.72	0.168	0.765	0.72	0.742	0.776	TRUE
	0.832	0.28	0.796	0.832	0.814	0.776	FALSE
Weighted Avg.	0.784	0.232	0.783	0.784	0.783	0.776	

E

Confusion Matrix

b | \leftarrow classified as $70 \quad a = TRUE$

Confusion Matrix

Appendix F

This appendix contains the results of Round 2B. The full dataset with the reduced 20 attributes is classified with ten-folds cross-validation.

MLP: Multilayer Perceptron

Summary					Confus	ion Mat	trix
Correctly Classi Incorrectly Class Kappa statistic Mean absolute of Root mean squa Relative absolut Root relative sq Total Number of Detailed Acc	sified Instan error ared error te error uared error f Instances	nces 137 0.58 0.27 0.38 55.7 77.0 670	6~%	% 	20 69 a =	classified = TRUE = FALSE	
Weighted Avg.	TP Rate 0.761 0.822 0.796	FP Rate 0.178 0.239 0.213	Precision 0.764 0.819 0.795	Recall 0.761 0.822 0.796	F-Measure 0.763 0.82 0.795	ROC Area 0.87 0.87 0.87	Class TRUE FALSE

SMO: Suport Vector Machine

Summary

Confusion Matrix

Correctly Classified Instances	540	80.60~%			
Incorrectly Classified Instances	130	19.40~%			
Kappa statistic	0.60			h	\leftarrow classified as
Mean absolute error	0.19		<u>a</u>		
Root mean squared error	0.44		220		a = TRUE
Relative absolute error	39.54~%		61	320	$\mathbf{b} = \mathbf{FALSE}$
Root relative squared error	88.94~%				
Total Number of Instances	670				

Detailed Accuracy By Class

	TP Rate	FP Rate	Precision	Recall	F-Measure	ROC	Class
						Area	
	0.761	0.16	0.783	0.761	0.772	0.801	TRUE
	0.84	0.239	0.823	0.84	0.831	0.801	FALSE
Weighted Avg.	0.806	0.205	0.805	0.806	0.806	0.801	

Logistic: Logistic Regression

Summary

Correctly Classified Instances	542	80.90~%			
Incorrectly Classified Instances	128	19.10~%			
Kappa statistic	0.61		а	b	\leftarrow classified as
Mean absolute error	0.26			~	
Root mean squared error	0.37		220	69	a = TRUE
Relative absolute error	53.81~%		59	322	b = FALSE
Root relative squared error	75.00~%				
Total Number of Instances	670				

Detailed Accuracy By Class

	TP Rate	FP Rate	Precision	Recall	F-Measure	ROC	Class
						Area	
	0.761	0.155	0.789	0.761	0.775	0.881	TRUE
	0.845	0.239	0.824	0.845	0.834	0.881	FALSE
Weighted Avg.	0.809	0.203	0.808	0.809	0.809	0.881	

k-NN: Nearest Neighbor

Summary

Correctly Classified Instances Incorrectly Classified Instances	$530 \\ 140 \\ 0.57$	$\begin{array}{c} 79.10 \ \% \\ 20.90 \ \% \end{array}$			
Kappa statistic Mean absolute error	$0.57 \\ 0.21$		a		\leftarrow classified as
Root mean squared error	0.46		$211 \\ 62$	78 310	a = TRUE $b = FALSE$
Relative absolute error	42.59 %		02	515	0 – MLDL
Root relative squared error Total Number of Instances	92.30 % 670				

Detailed Accuracy By Class

	TP Rate	FP Rate	Precision	Recall	F-Measure	ROC	Class
						Area	
	0.73	0.163	0.773	0.73	0.751	0.784	TRUE
	0.837	0.27	0.804	0.837	0.82	0.784	FALSE
Weighted Avg.	0.791	0.224	0.79	0.791	0.79	0.784	

Confusion Matrix

Confusion Matrix