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# A Machine Learning Approach to Diagnosis of Parkinson's Disease

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## Abstract

I will investigate applications of machine learning algorithms to medical data, adaptations of differences in data collection, and the use of ensemble techniques.

Focusing on the binary classification problem of Parkinson's Disease (PD) diagnosis, I will apply machine learning algorithms to a dataset consisting of voice recordings from healthy and PD subjects. Specifically, I will use Artificial Neural Networks, Support Vector Machines, and an Ensemble Learning algorithm to reproduce results from [MS12] and [GM09].

Next, I will adapt a regression dataset of PD recordings and combine it with the binary classification dataset, testing various techniques to consolidate the data including treating the regression data as unlabeled data in a semi-supervised learning approach. I will determine the performance of the above algorithms on this consolidated dataset.

Performance of algorithms will be evaluated using k-fold cross validation and a confusion matrix. Specificity and sensitivity will be calculated, as these are of particular importance in medical diagnosis. I will also determine accuracy, precision, recall, and F-score.

The expands on past related work, which has used either a regression dataset alone to predict a Unified Parkinson's Disease Rating Scale score for PD patients, or a classification dataset to determine healthy or PD diagnosis. The datasets have not been combined, and the regression set has not been used to contribute to evaluation of healthy subjects.

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## Chapter 1

# Introduction

Parkinson's Disease (PD) is a degenerative neurological disorder marked by decreased dopamine levels in the brain. It manifests itself through a deterioration of movement, including the presence of tremors and stiffness. There is commonly a marked effect on speech, including dysarthria (difficulty articulating sounds), hypophonia (lowered volume), and monotone (reduced pitch range). Additionally, cognitive impairments and changes in mood can occur, and risk of dementia is increased. PD affects about 1-2% of people over 60.

Traditional diagnosis of Parkinson's Disease involves a clinician taking a neurological history of the patient and observing motor skills in various situations. Since there is no definitive laboratory test to diagnose PD, diagnosis is often difficult, particularly in the early stages when motor effects are not yet severe. Monitoring progression of the disease over time requires repeated clinic visits by the patient. There is no cure, but pharmacological treatment to manage the condition includes dopaminergic drugs.

Diagnosis is clearly a difficulty in PD management, and an effective screening process, particularly one that doesn't require a clinic visit, would be beneficial. Since PD patients exhibit characteristic vocal features, voice recordings are a useful and noninvasive tool for

diagnosis. If machine learning algorithms could be applied to a voice recording dataset to accurately diagnosis PD, this would be an effective screening step prior to an appointment with a clinician.

Actually, this sort of binary classification problem is common in many areas of medical diagnosis, and techniques that work well within one domain are likely to be applicable to others. Based on prior work with PD as well as other disorders, support vector machines and artificial neural networks are ideal candidates for this type of problem. Another option for exploration is AdaBoost, an ensemble approach which combines learners in a way that minimizes their weaknesses.

However, one complication that may exist in available medical data is an inconsistency across datasets. For example, multiple datasets for a given disorder often exist, collected from different sources and using slightly different features. Combining them in some effective way into a large, cohesive dataset would result in a more robust and well-trained learner.

Another complication is an occasional lack of labeled examples. If some amount of labeled data exists, a semi-supervised learning algorithm might be applied, which uses a combination of labeled and unlabeled data. One approach is Expectation-Maximization (EM) which, given a model, alternates between making predictions about unlabeled data and recomputing the parameters of the model until convergence.

A successful machine learning approach to binary classification would be applicable to many types of medical diagnosis. Incorporation of semi-supervised learning with EM would result in a more robust system that is relevant to realistic medical datasets. Within the context of PD, this combination of techniques produces a useful screening step for a more frictionless diagnosis experience.



## Chapter 2

# Background

Many medical decision-making questions can be reduced to binary classification problems, making medical data an ideal domain for several machine learning techniques. A few of the most relevant algorithms, based on past work in this domain, are discussed below. Their applications to medical data in general, reviewed below, share many similarities with the specific case of Parkinson’s diagnosis.

### 2.1 Machine Learning techniques

#### 2.1.1 Artificial Neural Networks

Artificial neurons were first proposed in 1943. Motivated by biological neurons, the artificial neuron received several weighted inputs and produced an output, based on some threshold [MP43]. The perceptron model built on this early work, adding a learning rule to improve the performance of the neural network [Ros58]. However, the perceptron model was severely limited, unable to solve non-linearly separable functions such as XOR [MP69]. Eventually, multilayer perceptrons were developed to address the original perceptron model’s shortcomings.

Artificial neural nets have been widely used across a myriad of applications. For exam-

ple, a checkers-playing program used neural networks to train a player [CF01]. The board was encoded as a vector of available board positions, with values assigned based on whether the square was empty, taken by a regular piece, or taken by a King. The neural network consisted of three hidden layers. The first hidden layer completed spatial preprocessing, representing each subsquare of the board as a node, for a total of 91 nodes. The second and third hidden layers had 40 and 10 nodes, respectively. The network outputted a value between -1 and 1, representing the goodness of the board from the current player's perspective. The weights for the network were initially specified through a uniform sample, and several networks played against one another. The winners were declared 'parents', and they generated 'offspring networks' with weights varied by a parameter vector. The process was repeated for many generations to produce an ideal neural network.

### 2.1.2 Support Vector Machines

Support vector machines are binary classifiers that can be applied to linearly separable datasets. They separate data into classes using a hyperplane. SVMs can also be used non-linearly by mapping the data to a higher-dimensional space, thus making the data separable. This mapping is done by a kernel function. SVMs perform well with large feature spaces, as long as the data is separable with a wide margin. They also do well with sparse datasets, as in text classification [Joa98].

In the absence of large amounts of labeled data, pool-based active learning can be utilized with SVMs [TK01]. The learning algorithm has access to a pool of unlabeled data, and is able to choose a subset of that pool to use as training data. The learner chooses pool data to use such that the data minimizes the size of the learner's set of hypotheses, and brings it closer to a single hyperplane. This approach allows for using less labeled data.

### **2.1.3 Ensemble Learners**

Ensemble learners combine different machine learning algorithms. There is no one algorithm that always performs well on all domains, and ensemble learners are a way of combining the advantages of different learners. A good ensemble algorithm will be made up of diverse base learners that have varied strengths. The different learners can be combined in a number of ways. They can work in parallel on all of the inputs, and their outputs can be combined in some way. Alternatively, a multistage combination will train the base learners on different subsets of the input data. For example, the AdaBoost algorithm first trains an initial learner, and then trains subsequent learners on data that the first learner misclassifies. This way, the weaknesses of each base-learner are made up for by the next learner [FS95].

## **2.2 Applications to Medical Data**

Medical diagnosis presents an ideal domain for machine learning algorithms. A large part of diagnosis falls under pattern recognition, based on large amounts of data, and ML algorithms are well-suited to this task. For an algorithm to be effective in this domain, it needs to be able to handle noisy and missing data, rely on relatively few medical tests, and complement the role of physicians [Kon01]. Machine learning algorithms have been applied to a variety of medical data, some examples of which are outlined below.

### **2.2.1 Self-reported input**

Some diagnoses rely largely on patient-reported information, rather than biological tests. A prime example of this is diagnosis of mental disorders, which is based on how a patient's symptoms compare to criteria outlined in the Diagnostic and Statistical Manual of Mental Disorders. Symptoms are determined through consultation with a mental health professional, and are largely reported by the patient. Automated systems have been proposed

which will produce a diagnosis based on user-reported information [YC96].

### **2.2.2 Clinical Decision Support Systems**

Clinical decision support systems help healthcare professionals make diagnosis decisions based on patient data. These systems can be rule-based, in which case they are created with a knowledge base and a set of rules. Alternatively, they can utilize machine learning to learn from past data and recognize patterns. Several such systems have been proposed, including a statistical approach to diagnosing digestive disorders based on an electronically-administered interview of the patient [SK84]. However, the use is not yet widespread, in part due to lack of data availability and to limited adoption of uniform computer systems [Gre07].

### **2.2.3 EEG and EKG data**

Recordings of electrical activity in the body can be used to diagnose a variety of disorders. Electroencephalograms (EEGs) are recordings from the brain and contain a wealth of features that can be used by machine learning algorithms. A classification algorithm using EEGs was able to diagnose Alzheimer’s Disease with 86.05% accuracy [Pod12].

Electrocardiograms (EKGs) are often used to detect arrhythmia, which is any abnormality of the heartbeat. They can be indicative of heart disease and other conditions. In 1989, a model was derived from the Cleveland Clinic heart disease data set and compared it to CADENZA, a Bayesian algorithm. Both models were found to overpredict heart disease, though this occurred more with CADENZA [Det89].

After that, a novel machine learning approach to diagnosing and classifying cardiac arrhythmia was presented, called the VF15 algorithm. It used a genetic algorithm to learn feature weights. Then, each feature voted on a class prediction. The algorithm had a 62%

accuracy on this task and was found to outperform Naive Bayes [GAD98].

Another study collected data on ischemic heart disease, including signs and symptoms, EKG, and scintigraphy. Several algorithms were applied, including Naive Bayes, neural networks, k-nearest neighbors, and two decision tree algorithms. These were compared to clinicians diagnoses. Naive Bayes had the best sensitivity/recall, whereas clinicians, followed by neural nets, had the highest specificity [KKG<sup>+</sup>99].

More recently, a group compared various machine learning algorithms for arrhythmia diagnosis based on EKG data, with an emphasis on minimizing false positives and dealing with noisy data. They used the UCI Machine Learning Repository Arrhythmia dataset, and highlighted the need to improve on VF15s 62% accuracy. They evaluated a Bayesian artificial neural network classifier as compared to Naive Bayes, decision trees, logistic regression, and neural networks [GMCL05].

## 2.3 Diagnosing Parkinson’s Disease

Speech tests can be used for monitoring Parkinsons disease, due to vocal impairment being a common symptom and early indicator. Using an at-home recording device, such as one developed by Intel for PD telemonitoring, can conveniently allow PD patients’ health to be monitored remotely. Specified voice recordings can be passed through signal processing algorithms and a classification and regression tree to predict a rating on the unified PD rating scale [TLMR10].

Another study described a weakly supervised multiple instance learning approach to detecting symptoms of Parkinsons Disease. This approach addressed the issue of self-reporting resulting in inaccurate or incomplete data [DAITH12]. Their algorithm learned to localize symptoms to approximate, rather than exact, time ranges, making it suitable for the sparse data that may result from incomplete reporting.

Gil and Johnson used a multilayer network with one hidden layer and an output layer that output healthy or PD. The inputs were passed through a sigmoidal activation function, and gradient descent backpropagation was used to modify the weights. They achieved a classification accuracy of 92.31%. They also trained an SVM using the sequential minimal optimization (SMO) algorithm. SMO speeds up training of SVMs, particularly those with non-linear kernel functions (Platt 1998), using a divide and conquer approach. Gil and Johnson used a linear kernel with 91.79% accuracy, and a Pearson VII function kernel, with accuracy of 93.33% [GM09].

Mandal and Sairam also used a neural network with a sigmoidal activation function. They modified weights using backpropagation with dynamic learning rate and momentum, and achieved an accuracy of 97.6471%. They also used SVM with a linear kernel and obtained an accuracy of 97.6471% [MS12].

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