

## Case Study Application for C-Support Vector Classification: The Estimation of MS Subgroup Classification with Selected Kernels and Parameters

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**Abstract.** The study has classified the subgroups of Multiple Sclerosis (MS) using Support Vector Machines (SVM). C- Support Vector Classifier (C-SVC) algorithm, one of the SVM classifiers of multi class, has been utilized for the classification of MS subgroups. For this purpose, 120 MS patients (76 RRMS, 38 SPMS, 6 PPMS patients) have been included in the study. Through Magnetic Resonance Imaging (MRI), the number of lesion diameter and Expanded Disability Status Scale (EDSS) data are applied through C-SVC. Lesion data has been obtained from three separate regions of the brain which are brain stem, periventricular corpus callosum and upper cervical region. By applying the data onto Radial Basis Function kernel (RBF), Polynomial kernel, Sigmoid kernel and Linear kernel, four of the kernel types of C-SVC algorithm, the accuracy rates of MS subgroups classification and the computation time during the training procedure are computed and compared. By adding EDSS score into the dataset, the classification achievement rate has increased in all the kernel types based on the analyses conducted. Having applied C-SVC on MS subgroups, classification achievement of MS subgroups, namely that of RRMS, SPMS and PPMS has been measured.

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### 1. Introduction

Multiple sclerosis (MS) is a disorder of the Central Nervous System (CNS) that affects the brain and spinal cord and is accompanied by inflammatory, demyelinating and axonal damage. CNS transmits electrical messages in different parts of our body along the nerves. Such messages have control over all of our movements, both voluntary and involuntary. MS

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disorder, however, disrupts the proper transmission of these messages. The disorder causes neuronal dysfunction, bringing about the likelihood of long-term permanent damages. At the initial stage, a specific reason for the disorder is not known, thus it could be improbable to prevent and treat the disorder. In our day there exist no treatments that provide cure for MS. Yet, in the last two decades, several treatments have been developed and put into use, which reduce frequency of attacks, increase in the number of the plaques traced in MRI and permanent impairment in the long-run [22].

MS is categorized into 4 different clinical subtypes based on the attacks and existence of progression. The most frequently-seen one, called Relapsing Remitting MS (RRMS), is seen in 70% of the patients. Patients experience neurological loss during the attacks of RRMS based on the localization of the lesions. After the attack is over, the patients go through the remission phase and findings followed during the attack period are lost. About 50% of the patients of RRMS go under a progressive stage in which the attacks do not have remission in a period of 10 years. This form is called Secondary Progressive MS (SPMS). The patients afflicted by this form are observed to have accumulated neurological impairment over the years. In some patients, attacks and progression are seen in the course starting with the beginning phase of the disorder. This is a form called Relapsing Progressive MS (RPMS) in which the impairment gains a permanent state following each attack, and an accumulation of impairment is seen during the attack. In the rare form of MS, Primary Progressive MS (PPMS), attacks are not seen in the picture but similar to the case of SPMS, accumulation of neurological impairment is observed in the patients over the years. PPMS is seen in 10 - 15% of the patients, and its diagnosis is of critical importance since it is the form of MS that has the highest resistance to treatment [17, 21, 23, 26–28]. It is important that the subtypes of the disorder and their likelihood of causing disability in the future be identified in the early phase. It is also important to identify and compare the MRI lesion characteristics of the groups corresponding as well as the lesion load so that the treatment regimen can be figured out at an early stage and indicate the success of follow-up.

For the clinical diagnosis of MS disorder and its subgroups, lesion size received from the MR images of the patients taken on regular basis throughout the years and EDSS score based on McDonald criteria are used in this study [16]. The dataset is made up of data belonging to 120 MS patients.

Performance analysis of the MS subgroups among these patients based on different kernel types of the SVM machine learning is performed in this study. MS disorder is seen more frequently with the following characteristics: young adults aged between 20 and 50, Caucasians, people living in mild and cold climate zones, those whose family members have MS, people with a high profile in terms of social, cultural and economic aspects [21]. Data belonging to the patients and age interval table are presented in Table 1. Including 120 MS patients (aged between 20 and 55), the study performs the identification of RRMS, SPMS and PPMS. MR images of the patients taken from the last three years have been examined focusing on three regions, which are the brain stem, corpus callosum and upper part of the neck. At least 3 years have passed between the first MRI and second MRI, and at least 8 years have passed between the second and third MRI. The outcomes of the MS patients have been identified (associated) through EDSS scores by a neurologist using MRI results. While identifying the subgroups of the

disorder, MRI data has been used by examining the EDSS scores (the number of lesion diameter for three different regions identified was taken as a parameter). The multi class library of LibSVM of SVM supervised algorithm, one of the machine learning algorithms, has been used in this study for the diagnosis of the disorder for RRMS, SPMS and PPMS. Two different sets of data have been studied for the multi class SVM applying Radial Basis Function, Polynomial, Sigmoid and Linear kernels. For the first data set, using the number of each lesion diameter (width) for three different parts separately, a matrix of size  $120 \times 227$  has been obtained. The second data set yields a matrix of  $120 \times 228$  with EDSS scores and MRI data of 120 patients (the number of lesion diameter / width for three different regions separately). So as to see the significance of the parameter, EDSS score has been removed from the second dataset.

C-SVC (C-Support Vector Classification) is a multiclass Support Vector Machine algorithm method. Data has been applied onto methods such as Radial Basis Function (RBF) kernel, Polynomial kernel, Sigmoid kernel and Linear kernel which are considered as C-SVC algorithms [18]. According to C-SVC kernels experimental results have been given by two aspects that are k-fold cross validation and computation time. In the meantime, performance evaluation has been made through k-fold cross validation ( $k = 10$ ). The computation time during the classification of the C-SVC kernel (Radial Basis Function kernel, Polynomial kernel, Sigmoid kernel, Linear kernel) methods have been provided in a comparative fashion. Please see Table 2 for a brief review of relevant literature.

Table 1: Age Intervals Based on the Gender of the Patients in the Dataset.

Age	18-25	26-30	31-35	36-40	41-45	46-50	51-65
Female	10	22	17	16	9	8	6
Male	6	11	4	6	4	0	1

Table 2: Literature Review on MS.

Researchers	Experiments	Experimental Results
Gutermana, <i>et al</i> [14]	Data concerning 100 MS patients and control group with 100 individuals (brainstem trigeminal) evoked potential data was applied to Multi-Layer Perceptron Probabilistic Neural Network and Kohonen's Learning Method diagnosing Multiple Sclerosis. The performance of the neural networks based classifiers is compared with that of the human experts and the Bayes classifier.	Kohonen Learning: <b>91.91%</b>  Bayesian Learning: <b>96.01%</b>

Table 2: Literature Review on MS.

Researchers	Experiments	Experimental Results
Bergmasci, <i>et al.</i> [6]	Bayesian method was used for the early prediction of Multiple Sclerosis (MS) in the long-term. Bayesian Risk Estimation score (Brems) was calculated for each patient in the first year of the disorder by means of the Bayesian model used here.	<b>Brems Value:</b> SP within 10 years compared with 1087 progression 158 free patients: $p < 0.0001$ SP risk in the whole cohort: $p < .0001$ Subgroup of 535 patients who had never been treated with immune therapies: $p < 0.000001$
Hirst, <i>et al.</i> [17]	Analyses of t-test and chi square tests were compared in order to predict the natural course of MS disorder in MS patients and to understand the long term consequences.	Mean EDSS score in 1985: 5.15 (SD 2.7, range 0-9.5) and 8.01 (SD 2.6, range 0-10)  Mean worsening of EDSS scores in surviving patients was +3.02 EDSS points,  61.4% of patients with EDSS 3.5-5.5 and 82.2% of those with an EDSS of $\leq 3$ in 1985 had an EDSS of $\geq 6$ after 20 years.  Lower baseline EDSS scores ( $p < 0.0001$ ), higher pyramidal functional system score ( $p = 0.02$ ) and a greater number of functional systems involved ( $p = 0.001$ )  Of those with benign disease in 1985, only 19% remained benign after 20 years of follow-up 12.6% of patients had minimal disability after at least 20 years after their disease onset and 14% of patients failed to worsen by $\geq 1$ EDSS point.

Table 2: Literature Review on MS.

Researchers	Experiments	Experimental Results
Shahlaei, <i>et al.</i> [33]	Using data belonging to MS patients, Principal Components Regression (PCR), Principal Components Artificial Neural Networks (PC-YSA) and Principal Components least squares support vector machine (PC-LS-SVM) were examined with regression analyses.	<b>PC-ANN : 93.7%</b> <b>PC-SVM : 95.6%</b>
Karaca, <i>et al.</i> [28]	MRI results are evaluated by examining probability of Mean Echo, White matter, Gray Matter, Ventricular CSF, External CS. Diagnosis of RRMS, SPMS and PPMS disorder according to MRI1, MRI2, MRI3 were identified by decision tree method. MRI results were evaluated by examining the lesion counts and EDSS results according to Chi-Square Test of Independence method.	<b>Decision Tree Accuracy Rate:</b> MRI1:89.36% MRI2: 85.10% MRI3:93.61%  <b>The Chi-Square Test of Independence Results:</b> MRI1:0.292 MRI2:0.374 MRI3:0.097
Karaca <i>et al.</i> [24]	120 MS patients and 19 healthy individuals make up the dataset. Lesion data from MR images and EDSS scores of the patients were formed as the dataset for the study. Dataset 1 included EDSS scores and lesion data while Dataset 2 only had lesion data. By applying Learning Vector Quantization, Feed Forward Back Propagation (FFBP) and Radial Basis Function (RBF) algorithms in Artificial Neural Networks on Dataset1 and Dataset 2, the accuracy rate for classifying the subgroups of MS was obtained. As a result of the study, the significance of EDSS in terms of accuracy in classifying the subgroups of MS was revealed.	<b>DataSet1:</b> LVQ : 89% FFBP: 92.5% RBF : 98.9%  <b>DataSet2:</b> LVQ : 91% FFBP : 96.75% RBF : 99.9%

## 2. Material and Methods

### 2.1. Patient Details

In this study, monitored by Hacettepe University Faculty of Medicine, Departments of Neurology and Radiology and Primary MR Image center, 120 patients with a definite MS diagnosis based on the McDonald criteria with RRMS, SPMS or PPMS were enrolled. All patients were between the ages of 20 and 55.

Level of disability in patients was determined by using the Extended Disability Status Scale (EDSS). MRI is obtained with a 1.5 Tesla device (Magnetom, Siemens Medical Systems, Erlangen, Germany). Lesions on T2-weighted turbo spin-echo (TSE) sequences were counted in metric units. The brain stem, corpus callosum-periventricular region, including the upper cervical lesions in the three regions were included in the information. Magnetic resonance read for three regions lesion in the years to the information changes (number of increments/reductions in size) were compared based on the EDSS scores of years and changes within the clinical diagnostics were compared with MS. While observing the duration of the disease, there were minimum three years between the first and second MRI scans and there were maximum 8 or 10 years between the second and third and MRI scans. The outcomes of the MS patients have been identified (associated) through EDSS scores by a neurologist using MRI results [23, 24, 26, 27].

### 2.2. Expanded Disability Status Scale

Expanded Disability Scale (EDSS) is partly based on the measurements of eight regions known as functional system in the central nervous system. First, the scale measures the temporary numbness in the face or fingers or degrees of impairment such as visual impairment. And then, by using walking distance, it measures the disability in terms of mobility [26].

Functional systems measured with EDSS:

- (i) Pyramidal: voluntary movements
- (ii) Brain stem: eye movements, senses, face movements, functions like swallowing
- (iii) Vision
- (iv) Brain: memory, concentration, temperament
- (v) Cerebellum: balance and coordination of movements
- (vi) Sense
- (vii) Bowel and bladder
- (viii) Others: including fatigue

Table 3: Description of EDSS Scores. [6, 12, 26]

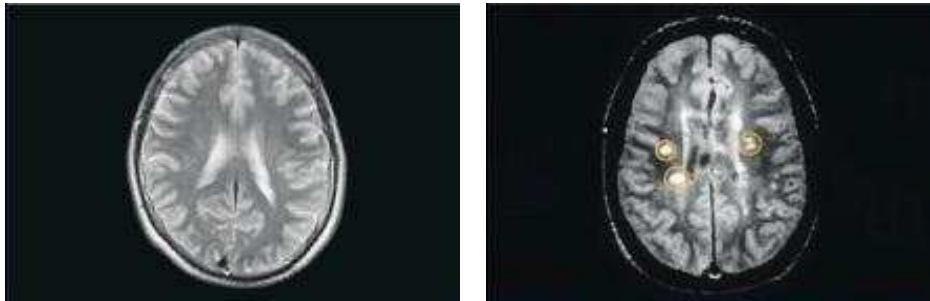
Score	Description
1.0	No disability, minimal signs in one FS
1.5	No disability, minimal signs in more than one FS
2.0	Minimal disability in one FS
2.5	Mild disability in one FS or minimal disability in two FS
3.0	Moderate disability in one FS, or mild disability in three or four FS. No impairment to walking
3.5	Moderate disability in one FS and more than minimal disability several others. No impairment to walking
4.0	Significant disability but self-sufficient and up and about some 12 hours a day. Able to walk without aid or rest for 500m
4.5	Significant disability but up and about much of the day, able to work a full day, may otherwise have some limitation of full activity or require minimal assistance. Able to walk without aid or rest for 300m
5.0	Disability severe enough to impair full daily activities and ability to work a full day without special provisions. Able to walk without aid or rest for 200m
5.5	Disability severe enough to preclude full daily activities. Able to walk without aid or rest for 100m
6.0	Requires a walking aid - cane, crutch, etc - to walk about 100m with or without resting
6.5	Requires two walking aids - pair of canes, crutches, etc - to walk about 20m without resting
7.0	Unable to walk beyond approximately 5m even with aid. Essentially restricted to wheelchair; though wheels self in standard wheelchair and transfers alone. Up and about in wheelchair some 12 hours a day
7.5	Unable to take more than a few steps. Restricted to wheelchair and may need aid in transferring. Can wheel self but cannot carry on in standard wheelchair for a full day and may require a motorized wheelchair
8.0	Essentially restricted to bed or chair or pushed in wheelchair. May be out of bed itself much of the day. Retains many self-care functions. Generally has effective use of arms
8.5	Essentially restricted to bed much of day. Has some effective use of arms retains some self-care functions
9.0	Confined to bed. Can still communicate and eat.
9.5	Confined to bed and totally dependent. Unable to communicate or eat/swallow
10.0	Death due to MS [12, 26]

According to Table 3, these systems are rated according to the degree of the impairment. These

degrees can be 0 for normal condition and may rise up to 5 or 6 for conditions where the impairment is highest. By adding the movement and daily life limitations to these functional system degrees, we defined the 20 steps of EDSS [21]. The EDSS quantifies disability in eight Functional Systems (FS) and allows neurologists to assign a Functional System Score (FSS) in each of these data are applied by Tanagra software [12, 26].

### 2.3. Magnetic Resonance Imaging (MRI)

MR scanner utilizes strong magnetic fields to yield the view of the brain and spinal cord. An MR image can reveal inflammatory or damaged tissue regions in the central nervous system [16, 22]. Figure 1a presents an MR image of a healthy individual while Figure 1b shows the MR image of an MS patient [23, 26]. Lesion numbers in the MR image of patients taken by a radiologist at regular intervals have been used for the study.



(a) Healthy Patient [23, 26]

(b) MS Patient [23, 26]

Figure 1: MR Images of Patient's Brains

In this study, we used the number of lesions according to the locations of the brain, and EDSS score for modeling in Table 4.

Table 4: Feature Extracted from MR Images and Representation of Classes and EDSS.

Feature Explanation	Explanation
EDSS	ranges between 1 and 10
RRMS	Integer Number (0)
PPMS	Integer Number (-1)
SPMS	Integer Number (1)
Lesions	the number of lesion diameter that has formed in three different parts of the brain

The data in the study has been taken at regular intervals, as stated in Table 4 and Table 5. Using two different training sets, as stated in Table 5. The data of the study is comprised of lesion diameter from MR images as well as EDSS scores. According to Table 4, this study



has 3 classes (RRMS, PPMS, SPMS), and the data has been applied on Radial Basis Function, Polynomial, Sigmoid and Linear kernels.

Table 5: Training Set Matrix Dimensions

	Vector Size
Training Set 1	120x227
Training Set 2	120x228

### 2.4. Support Vector Machines

SVM is a modern classifier which is able to make very good generalizations structuring linear classification boundaries in multidimensional space through kernel [5, 9, 25, 30, 31, 36]. In this study, the subgroups of the disorder, namely RRMS, SPMS, and PPMS as well as classifier performance estimations for MS subgroups has been obtained on different kernels, using the LIBSVM implementation of the SVM algorithm [7]. SVM design is split by hyperplane [31] which represents the decision limits; the observations which define the boundaries are called "support vectors". Class estimation is made based on these decision limits. In SVM, a maximum level of accuracy is achieved in the estimation of class prediction of a new set of data formed through the use of optimum decision limit from training data. When class representation belonging to the data is not divided in linear fashion, the formation of decision limit on extreme plane between the classes is an optimization problem. For the solution of such a problem, it is necessary to identify the function between two points as  $(x, f(x))$  using the variables in the data set, as seen here:

$$f_0(x) \tag{1}$$

$$f_i(x) \leq 0, i = 1, \dots, m \tag{2}$$

$$h_i(x) = 0, i = 1, \dots, p \tag{3}$$

If the classes cannot be classified in a linear fashion, the optimization problem in SVM is placed to the new space through non-linear based functions. The length of the new space is bigger than that of the initial space [13]. The complexity of it is done through a learning method that is not related to the number of dimension. While finding the best hyperplane, complexity in the optimization problem is written based on the sample number (120). In this way, cross checking has been done while transferring from base functions to the kernel functions. The point that is the least based on  $w_0$  values or the biggest point based on  $a^T \geq 0$  values is found [13] ( $\alpha$  and  $w$  will be defined shortly).

$x \in R^n$  represents optimization variable,  $f_0 : R^n \rightarrow R$  is the cost function. It represents the constant functions that are not equal.  $h_i : R^n \rightarrow R$  represents the cost function that is equal [29].

$$p^* = \inf f_0(x) | f_i(x) \leq 0, i = 1, \dots, m, h_i(x) = 0, i = 1, \dots, p \tag{4}$$

Based on equation 4, the optimum value is represented. If the problem is infeasible, it is  $p^* = \infty$ , If the problem is infinite, then it is  $p^* = -\infty$  [15].

$x \in R^n$  means variable, D means domain,  $p^*$  means optimum value [15]. Lagrangian;

$$L : R^n \times R^m \times R^p \rightarrow R, \text{ with } \text{dom}L = D \times R^m \times R^p, \tag{5}$$

With Lagrangian Dual Function the lowest level  $p^*$  is found.

$$g : R^m \times R^p \rightarrow R \tag{6}$$

$$g(\lambda, \nu) = \inf_{x \in D} L(x, \lambda, \nu) \tag{7}$$

$$= \inf_{x \in D} (f_0(x) + \sum_i^m \lambda_i f_i(x) + \sum_i^p \nu_i h_i(x)) \tag{8}$$

$$\min_{w,b,\xi} \frac{1}{2} w^T w + c \sum_{i=1}^l \xi_i \tag{9}$$

$\phi(x_i)$ ,  $x_i$  represents the data set in Training Set 1 and Training Set 2. For Training Set 1  $x$  is  $120 \times 227$ , and for Training Set 2  $x$  is  $120 \times 228$  sized matrix.  $c$  parameter represents our classes, and these are RRMS, SPMS and PPMS.  $w$  solves the dual problem in the data set with 3 classes through the equation 10 given below.

After having reached the solution with equation for the dual problem [25, 26], the optimum value is found for  $w$ ,

$$w = \sum_{i=1}^l y_i \alpha_i \phi(x_i) \tag{10}$$

through [13].

$$\text{sgn}(w^T \phi(x) + c) = \text{sgn}(\sum_{i=1}^l y_i \alpha_i K(x_i, x) + c) \tag{11}$$

the class determination of MS subgroups, the decision is given based on the function [7, 13].

Based on parameters, with Grid Search, direct learning is not done for class definers. The best defining performance is done based on cross-validation for this. The frequently used parameter for this is regularizer parameter. The performance evaluation for this study has been performed through multi-class [9].

A second important point in SVM is that the kernel use named as kernel trick in training data samples is in implicit nonlinear form. For instance, the form of data with its dot product taken (inner product) like the ones given in Figure 2 represents the kernel trick. In this way, it is possible to work by passing into different spaces in the data by setting up a linear model (right pane) instead of using nonlinear models (left pane). Data should be split initially by a simple hyperplane to be able to pass into different spaces. C-SVC algorithm was used, which is able to classify SVM data with multiple classes.

Kernel trick acts like a bridge from making transfer from linearity to non-linearity. In reality, it makes the big sized and non-linear input data like a linear algorithm. If the problem is not linear, learning can be done like a linear model by going to a new space through non-linear

\*Images from [2]

base functions. The linear model in the new space corresponds to the non-linear model in the initial space. This approach is used for classification [3, 11, 32, 34].

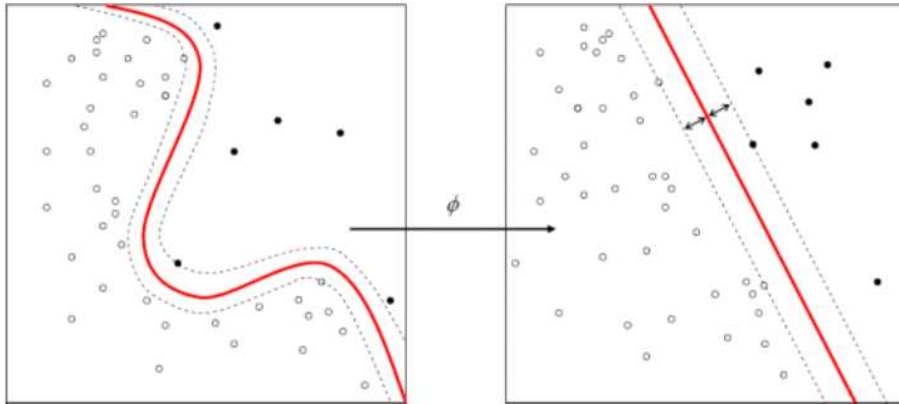


Figure 2: Demonstrating the Kernel Trick, from a Nonlinear Model in the Original Data Space (left pane) to a Linear Model in the Feature Space (right pane), via the implicit map  $\phi$ . [30]

$$L_p = 1/2 \|w\|^2 - \sum_{t=1}^N \alpha^t [r^t (w^T x^t + w_0) - 1] \tag{12}$$

$$L_p = 1/2 \|w\|^2 - \sum_{t=1}^N \alpha^t [r^t (w^T x^t + w_0) + \sum_{t=1}^N \alpha^t] \tag{13}$$

Through equation 13, Lagrange terms are added. Derivative is taken based on the parameters and reset to zero.

$$\partial L_p / \partial w = 0 \Rightarrow w = \sum_{t=1}^N \alpha^t r^t x^t \tag{14}$$

$$\partial L_p / \partial w = 0 \Rightarrow \sum_{t=1}^N \alpha^t r^t = 0 \tag{15}$$

In this case, the conjugate,

$$\begin{aligned} L_d &= 1/2 (w^T w) - w^t \sum_t \alpha^t r^t x^t - w_0 \sum_t \alpha^t r^t + \sum_t \alpha^t \\ &= -1/2 (w^t w) + \sum_t \alpha^t \\ &= -1/2 \sum_t \sum_s \alpha^t \alpha^s r^t r^s (x^t)^T x^s + \sum_t \alpha^t, \end{aligned} \tag{16}$$

is obtained as equation 17. The main idea in Kernel machines is to be able to write the inner product on base functions as a kernel function, which means  $K(x^t, x)$  on the initial inputs

[11, 20, 32, 34]. If there is a kernel function, there is no need to conjugate to the new space. In reality, there is a kernel function that corresponds to for each valid kernel but it is easier than using  $K(x^t, x)$  [11, 20, 32, 34].  $\phi(x^t)$  or  $\phi(x)$  values. In recent years, it has become a more common practice to keep the data sets in  $K$  kernels instead of  $\phi(x)$ . It is particular easier for keeping the  $N \times N$  matrix [3, 11, 20, 32, 34].

C-SVC represents two classes or multiple classes.  $C$  represents the number of classes. This study  $C$  has three classes which are, RRMS, SPMS and PPMS.

Accuracy rate estimation has been made by getting a class achievement as a result of applying the Radial Basis Function, Polynomial, Sigmoid, Linear, respectively. C-SVC kernel has been applied separately onto the dataset split by hyper plane, and the class achievement has been measured based on 10-fold cross validation. The two class training set is,  $X_i \in R^n, I = 1, \dots, l$  class labels are  $y_i \in [-1, 1]$  [10, 30, 31].

The classification optimization of C-SVC algorithm is; constraints are:

$$y_1 (w^T \Phi(x_1) + b) \geq 1 - \xi_i, \xi_i \geq 0, i = 1 \dots l \tag{17}$$

Definition of dual problem is as follows:

$$\min_{\alpha} \frac{1}{2} \alpha^T Q \alpha - e^T, 0 \leq \alpha_i \leq C, i = 1 \dots l \tag{18}$$

With constraints  $y^T \alpha = 0$ , where  $e$  is a vector of all ones  $C > 0$  is upper bound,  $Q$  is a  $l \times l$  positive semi define matrix  $Q_{ij} = y_i y_j K(x_t, x)$  and  $K(x_t, x) = \phi(x_t^T) \phi(x_j)$  is called the kernel function  $\phi$  transforms training vectors  $x_t$  into a higher dimensional space. In this study, there are two different data sets.

Decision function is;

$$\text{sgn} \left( \sum_{i=1}^l y_i \alpha_i K(x_t, x) + c \right) \tag{19}$$

$$K(x_t, x) = \exp(-\gamma \|x_t - x\|^2), \gamma > 0 \tag{20}$$

Laplacian Kernel and ANOVA Kernel [11] are equivalent of RBF kernel. Rational Quadratic Kernel is an alternative when we wish to use Gaussian. Multiquadric Kernel is non-positive defined kernel. Inverse Multiquadric Kernel [29] is appropriate for feature space in infinite dimension. Circular Kernel [32] and Spherical Kernel are appropriate for Geostatic applications, and Wave kernel [28] for semi-definite symmetrical positive data. Power Kernel [32] proves to be appropriate for positive defined conditional applications and Spline Kernel is appropriate for partial cubical polynomial data [30]. Log Kernel is used on images. Bessel Kernel fractional smoothness is also used. Cauchy Kernel [4] is used more in space with big dimension. Chi-Square Kernel [34] is used for applications that have to be positive defined that is conditional. Histogram Intersection Kernel is used for image recognition, Generalized Histogram Intersection Kernel [8] for the applications with wide contexts. A Bayesian Kernel is used in the prediction of protein-protein interaction data [1, 19]. For practices like system and cybernetics Wavelet kernel type is utilized [35]. While applying the MS data set on SVM algorithm,

optimization problem has arisen since the data set has multiple classes. It has been possible to overcome this situation by doing experimental applications on different kernel types. The optimized values of the parameters in the data set have been presented in Table 6. As a result of our experimental studies, the best classifying estimation in SVM has been yielded through Linear, RBF, Polynomial and Sigmoid Kernel on account of the nature of our data set. For this reason, these four main kernel types have been preferred [15].

Table 6: C-SVC Kernel Parameters

	Kernel Type			
	LINEAR	POLY	RBF	SIGMOID
<b>Degree(poly)</b>	1.00	2.00	1.00	3.00
<b>Gamma in kernel function (poly/rbf/sigmoid)</b>	0			
<b>Coef in kernel function</b>	0			
<b>Tolerance of termination criteria (eps)</b>	0.001			
<b>C (Complexity Cost)</b>	1			
<b>Compute probability estimates</b>	1			
<b>Use shrinking heuristics</b>	1			
<b>Data normalization</b>	1			

As to (Radial Basis Function) Kernel, in the study Gaussian kernel type has been used. Sigma ( $s$ ) plays a major role in the performance of the kernel.  $s$  value should be tuned with keen attention based on the available problem. If an extreme value is assigned for the sigma, the exponential value acts almost linearly and loses power for the big sized non-linear data [11, 15, 20]. Thus, there will be deprivation of function regulation, and the determination threshold will be sensitive for noise. Gaussian kernel has been chosen for the MS data set that is not classified linearly. For this reason, it has been deemed to be the appropriate kernel in our data set.  $s$  value is chosen to be 3.

The equation,

$$K(x^t, x) = \exp\left(\frac{-\|x^t - x\|^2}{2s^2}\right), \quad (21)$$

represents the global kernel.  $x^t$  determines the center and  $s$  determines the radius that has been formerly made constant. When the radius is large, hyperplane becomes more flat. The best results are found through cross validation. While adjusting the two upper parameters with cross validation, searching is done in two dimensions for all ( $c$  and  $s^2$ ) possible value duals.

Polynomial kernel is a kernel that is appropriate for the non-stationary data set. Like the data set used in our study, it is the appropriate kernel type for the normalized training data set [30]. For degree  $q$ ,

$$K(x^t, x) = (x^T x^t + 1)^q, \quad (22)$$

it is written as equation 25. And it is necessary that the  $q$  is predetermined. For th quadratic

kernel,  $q$  degree is chosen as 2; for  $q = 2$ :

$$\begin{aligned} K(x, y) &= (x^T y + 1)^2 \\ &= (x_1 y_1 + x_2 y_2 + 1)^2 \\ &= 1 + 2x_1 y_1 + 2x_2 y_2 + 2x_1 y_1 x_2 y_2 + x_1^2 y_1^2 + x_2^2 y_2^2 \end{aligned} \quad (23)$$

The kernel,

$$\phi(x) = (1, \sqrt{2x_1}, \sqrt{2x_2}, \sqrt{2x_1 x_2}, x_1^2, x_2^2)^T \quad (24)$$

corresponds to the inner product of the base function.

Sigmoid Kernel is known as the Multi-layered Perceptron kernel [20]. It is used as the activation function in artificial neurons. This kernel has been preferred since the MS data set is appropriate for multi-layer learning. In addition to this, it yields a good level of performance in practice although it is conditionally positive definite [3, 29]. Alpha value gets  $1/N$  value, where  $N$  is the number observations. For both our training and testing sets,  $N = 120$ .

$$K(x^t, x) = \tanh(\gamma \langle x^t, x \rangle + c) \quad (25)$$

$$K(x^t, x) = \tanh(\langle x \rangle + c) \quad (26)$$

ranges between -1 and +1.

Linear Kernel is the most simple kernel, and  $\langle x, y \rangle$  is given in the input.  $c$  has constant value. It is used generally in kernel algorithms [15].

$$K(x^t, x) = \langle x^t, x \rangle + c \quad (27)$$

$$K(x^t, x) = x + c \quad (28)$$

The main purpose of our study is to be able to define the subgroups of MS, namely RRMS, SPMS and PPMS through SVM algorithm. At the same time, the purpose is to show the accuracy of using multi kernel use in defining MS subgroups. SVM algorithm makes the class estimation of the data with classes more than two with higher accuracy. In this respect, this study is the first of its kind in literature since it has conducted the performance evaluation of C-SVC algorithm kernel types for the classification of MS subgroups on MS data set.

Many different practices have been applied for parameter values with Gamma in kernel function, Coef in kernel function, Tolerance of termination criteria, Complexity cost, Compute probability estimates, Use shrinking heuristics, and Data normalization. Applications have been made on RBF, Polynomial, Sigmoid, Linear kernel parameters between the lowest limit (0) and upper limit (4). Numeric parametric values provided in Table 6 are the ones optimized for the kernel types in the article. Some data set and parameters in LIBSVM library may reduce the iteration number in making the required calculations in classification [9]. The size of the MS data set in the study did not have a big size that would impede the making of the calculations.

### 3. Results

MR images and EDSS scores of have been utilized in our study while carrying out the diagnosis. MRI results are evaluated by examining the lesion counts and EDSS scores. LibSVM, multiclass library of SVM supervised algorithm which is one of the popular machine learning algorithms has been used for the diagnosis of the disorder regarding RRMS, SPMS and PPMS. The study has been conducted on two different data sets. In the first data set, EDSS score has been removed from the data set to emphasize the significance of the parameter. For this reason, it is a matrix of size  $120 \times 227$ . The second data set is a matrix of size  $120 \times 228$  with EDSS scores and MRI data of 120 patients (the number of lesion diameter/width for three different regions separately). The evaluation has been made by applying Radial Basis Function, Polynomial, Sigmoid and Linear kernel on two different datasets for multiclass SVM (C-Support Vector Classification) [10]. By using C-SVC [5, 25, 36] algorithm used for applications which SVM algorithm exceeds two classes, the performance classification analysis of MS subgroups (RRMS, SPMS, PPMS) has been performed. LibSVM library has been used for this purpose.

The verification has been made by 10-fold cross validation which is one method used in statistics. In addition to this, during the classification stage of C-SVC kernel methods (RBF kernel, Polynomial kernel, Sigmoid kernel, Linear kernel), computation time has been provided in a comparative fashion. Accuracy of the data has been validated and applied onto C-SVC algorithm kernel types separately. In addition, the verification of the accuracy rates has been made by 10-fold cross validation, which is one of the relevant statistical methods. In the end, while forming the classification of kernel types, the time elapsed and test achievement results have been compared with each other.

The results of our study have been obtained using Tanagra software, which is one of the data mining tools<sup>†</sup>. LibSVM library has been used for this purpose.

Table 7: Classification Achievement Rates and Classification Computation Time Based on Kernel Types for Training Set 1.

Kernel Type	Classification Rate	Computation Time (ms)
<b>Radial Basis Function</b>	99.658%	1217
<b>Polynomial</b>	99.633%	1154
<b>Sigmoid</b>	99.615%	1202
<b>Linear</b>	99.741%	858

<sup>†</sup><http://eric.univ-lyon2.fr/~ricco/tanagra/en/tanagra.html>

Table 8: Classification Achievement Rates and Classification Computation Time Based on Kernel Types for Training Set 2.

Kernel Type	Classification Rate	Computation Time (ms)
<b>Radial Basis Function</b>	99.675%	2137
<b>Polynomial</b>	99.635%	2122
<b>Sigmoid</b>	99.633%	2106
<b>Linear</b>	99.991%	1466

Based on analyses in Table 7, while forming the classification model belonging to kernel types applied onto the  $120 \times 227$  data set, working time elapsed and test achievement rate have been evaluated. The least classification duration belongs to the Linear kernel. When classification achievement rate is analyzed, the best performances are seen in Linear, RBF, Polynomial and Sigmoid kernel, respectively.

The Radial Basis Function (RBF) kernel has the highest duration of working time in the classification regarding  $120 \times 228$  data set given in Table 8. On the other hand, the lowest working time belongs to Linear kernel. When the performance result in classification is examined, the most successful kernel is Linear kernel, followed by RBF, Polynomial kernel, and Sigmoid kernel, respectively.

#### 4. Conclusions

The aim of this study is to be able to carry out a classification using C-SVC algorithm of SVM algorithm from the data set regarding the lesion counts in the lesion diameters obtained from the MR images of people with RRMS, SPMS and PPMS subgroups of MS. In addition, EDSS scores of each patient have been used. Based on the four different kernel types of C-SVC algorithm (Radial Basis Function, Polynomial, Sigmoid and Linear kernel) classification has been made.

Classification has been made based on the four different kernel types of C-SVC algorithm (Radial Basis Function, Polynomial, Sigmoid and Linear kernels). The number of lesion diameter taken from the MR images and EDSS scores of the MS patients make up the dataset. C-SVC algorithm has been applied onto kernel types separately, the achievements thereof have been verified and compared based on 10-fold cross validation, which is one of the relevant statistical methods.

The achievement of classification has been performed separately for kernel types. In each kernel time, the computation time has been calculated and the performance analysis has been carried out in accordance with 10-fold cross validation method. By adding EDSS score into the dataset, the classification achievement rate has increased in all the kernel types based on the analyses conducted. This rate has been found as 0.017% for RBF kernel, 0.002% for Polynomial kernel, 0.018% for Sigmoid kernel, and 0.25% for Linear kernel. As can be seen from this point, EDSS is a highly important factor in the diagnosis of MS. Sigmoid kernel has proved to be ineffective compared to other kernel types when the test achievement rate in



classification and the performance in classification are examined in both of the datasets. The reason for this is the  $\gamma$  and  $r$  parameters in Sigmoid kernel. Linear kernel type has been formed by adding the EDSS parameter into the most accurate kernel type dataset.

This study highlights the fact that it is possible to use C-SVC algorithm while doing classifications in static and non-static medical data. It has also been aimed that the study could provide a reference in the view that achievement rate can be received based on different kernel types. The findings the study have yielded, like the data such as lesion diameter, number etc. received from the MR images of the patients, EDSS and doctors' guidance were also elements that have been made use of.

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