Predictive Modeling of ECG Signal Using Spatial Topology Based Feature Space Transformation

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(ABSTRACT)

Biomedical signal classification has been frequently investigated by researchers in the past decades. Various classification systems have been proposed and evaluated by classification metrics. The essence of most methods, is to analyze test signals using reference models constructed based on a collection of healthy and abnormal signals. While system performances in terms of classifying signal samples increased significantly with modern machine learning algorithms such as recurrent neural network, there are one important factor in biomedical signal processing which is rarely studied in recent research. In this work, we go one step beyond the conventional methods and intend to predict potential upcoming abnormalities before their occurrence. The approach is to build a patient-specific model and identify minor deviations (e.g. yellow alarms) from the normal trend, which can be indicative of potential upcoming significant deviations (red alarms).

To facilitate a sound deviation analysis, a controlled spatial transformation is proposed to reshape the signal geometry in the feature space, such that the abnormality classes symmetrically surround the normal class. We applied the developed technique to Electrocardiogram (ECG) signals and the results confirm the utility of the proposed method in predicting upcoming heart abnormalities before their occurrence. For instance, the probability of a red alarm of specific abnormality class increases by 10% after observing a yellow alarm of the same type. This approach is general and has the potential to be applied to a wide range of physiological signals.

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To my family

Chapter 1

Introduction

1.1 Background and Motivation

In the last decades, mortalities caused by heart diseases has been increasing sharply, as a result of aging of population, chronic cardiovascular diseases, increasing life stress and continuously accelerating pace of life [1]. According to heart disease and stroke statistics, cardiac diseases are the most common cause of sudden cardiac death (SCD) which leads to 250 000 to 300 000 mortalities in J.S. every year which accounts for 14.7% of total deaths [2]. As World Health Organization reported, 31% of global deaths are related to cardiovascular diseases (CVDs) [3]. These facts fully reflect that heart diseases are threatening the general health of human beings. Meanwhile since SCD can occur, most likely, without warning and apparent symptoms in advance, the prediction and early recognition of cardiac disease are considered challenging [4]. Therefore it is of important significance to treat heart diseases timely by, early detection and prediction of CVDs. While most CVDs are accompanied with arrhythmia, accurate and timely recolonization of arrhythmia is a key factor for preventing the incidence of heart diseases effectively and determine definitive therapies for CVDs.

Electrocardiogram (ECG), recorded for the first time by Waller in 1887, contains abundant physiological and pathological information that reflects the heart rhythm and status of various parts of the heart [5]. ECG records signals generated by electrical activities of heart as a time series. As a noninvasive examination method, it is known to be highly reliable in reflecting functionality of heart. For this reason, ECG has become one of the most conventional technologies (ECG, clinical examination, radiation and ultrasonic inspection) in modernized hospitals and clinics, serving as an important reference for doctors diagnosis of heart diseases.

The traditional diagnosis based on ECG analysis are mainly accomplished manually by physicians. However, this becomes costly and impractical when continuous monitoring for CVDs is required. There are tremendous ECG records generated everyday, all demand for timely diagnosis and analysis. Facing with the severe shortage of physicians, automatic ECG classification system is introduced to generate real-time analysis result and provide additional information to physicians.

Automatic classification algorithms has been frequently investigated by researchers in the last decades. Whereas applying conventional classification algorithms on biomedical signals remains challenging, especially for applications of spontaneous disease detection.

A typical feature of cardiovascular disease is its spontaneous occurrence. This unpredictability is the principal cause of high mortality for patients with heart disease. An informative anticipation of abnormality would enable a therapeutic intervention ahead of mortal occurrence and thus minimize risk of mortality. Nevertheless, the majority of developed conventional ECG classification systems haven't focused on real-time or predicting performance.

Another important character of ECG waveform is its variability caused by distinct physical conditions of different individuals (i.e. gender, age, body-mass index etc.) [6] [7]. Con-

ventional classification algorithms fall short of generalization when applying on different patients' records [8]. Due to the inter-patient variation in ECG signals and the complexity of cardiac pathological information analysis, most of the existing ECG analysis software only serve as auxiliary devices for physicians. The final results of diagnosis still depend on manual labeling by cardiologists.

The automatic analysis of ECG signals incorporates a wide range of techniques. In this work, we focus on overcoming the two drawbacks of existing automatic ECG classification systems, which is patient-specificity and predicting capacity. This research aims at improving the inter-patient classification performance and predicting capacity of automatic analysis and recognition of arrhythmia with ECG signals, which has significance in efficient early detection of CVDs and reduce mortality rate of SCD.

1.2 ECG and Arrhythmia

Electrocardiogram is widely used to monitor the electrical activities of heart and assist diagnosing fatal cardiac disease. In order to design algorithms specifically for ECG analysis, it's important to develop a insightful perception of the functionality of heart and ECG waveform.

1.2.1 Characteristics of ECG signal

ECG signal reflects the periodical electric signals generated by heart. Fig.?? demonstrated the typical signal waveform for a cardiac cycle (i.e. a heartbeat), which is usually composed of three main waves; P wave, QRS complexes and T wave. These waves corresponds to different physiological activities of heart. P waves are generated by atrial depolarization

which standards for the process of pumping blood to ventricles. QRS complexes as the most significant electric activities is caused by ventricular contraction meaning the process of pumping blood to lungs and the rest of the human body. Finally, T waves is the result of ventricular repolarization recovering this process at the end of a cardiac cycle. Accurate detection and segmentation of each waves are necessary for ECG analysis. The waves are usually localized at the peak of waves, these peaks are also called fiducial peaks. By detecting the most significant peak within QRS complexes: R peak, automatic algorithms are able to draw boundaries between cardiac cycle. The interval between two R peaks is called RR interval, which is also the inverse of heart rate.

1.2.2 MIT-BIH Arrhythmia Database

Arrhythmia is related to various of morbid behavior of heart. Generally speaking, arrhythmias consist of two main categories: supraventricular and ventricular. Ventricular ectopic beats imply abnormal activities in the ventricles while Supraventricular ectopic beats are related to the atria. Both categories contains fatal abnormal beats which may lead to death. Therefore, in order to help researchers standardize the evaluation of works on ECG classifiers, Association for the Advancement of Medical Instrumentation (AAMI) [9] has proposed recommendations for reporting ECG classifier performance. According to these recommendations, MIT-BIH Arrhythmia Database (MITDB) is regarded as the standard database to train and test ECG classifiers in the last decades. MITDB is a public database which is available on Physionet.com [10] since 1997 [11]. There are 48 records collected from 47 individuals in the database. Each record contains two channel of ECG raw signals together with annotations composed of 16 types and R peak locations provided by cardiologists. The sample frequency of MITDB is 360Hz and signal frequency spans from 0.1 to 100 Hz.

Following the recommendation by AAMI, the original annotations of MITDB are further grouped into 4 major classes: class N(normal and bundle branch block beat types) class V(ventricular type), class S(supraventricular type) and class F(fusion of normal and ventricular types). The class Q which includes unclassified and paced beats are discarded due to the limit number of samples. Table.1.1 summarize the mapping from 16 original types in annotation to the standard 5 types recommended by AAMI:

Table 1.1: Mapping from 16 original types in annotation to the standard 5 types recommended by AAMI

Standard Types by AAMI	Original Types in MITDB Annotation
N	NOR, LBBB, RBBB, AE, NE
V	PVC, VE, VF
S	APC, AP, BAP, NP
F	VFN
Q	PACE, FPN, UN

1.3 Problem Statement

ECG signals, as a non-invasive method, are investigated broadly by researchers to design automated analysis and real-time monitoring systems. In the past decades, various ECG automatic analysis systems are proposed. As described in the last sections, there exists two main challenges: inter-patient variability and requirements for early detection and prediction.

For conventional classification system, the training data is composed of records collected from different patients with annotations per heartbeat. In order to unify the records from different patients, most of the algorithms using conventional classification algorithm concatenate heartbeats from different records and thus result in a pooled ECG dataset. Since the classification performance is measured based on generated labels for each sample, the classifiers are trained to improve the performance on pooled ECG data. While ECG signals

shares similar morphologies, the signals from different patients demonstrate considerable variance as shown in Fig.1.1. Ignoring this difference will lead to inconsistent classification performance between patients. Therefore it's of significant importance to adjust classifier configuration according to patient-specific characteristics.

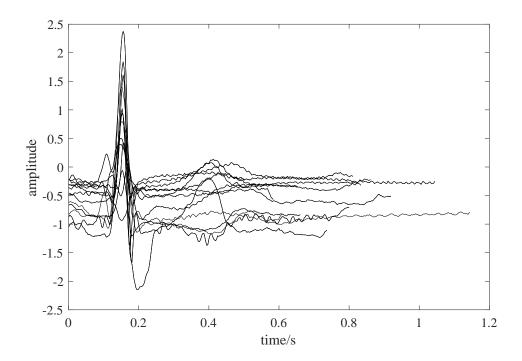


Figure 1.1: ECG signals of normal heartbeat from 15 different records reflect the inter-patient variability of ECG signal

In addition to the inter-patient variability, majority of ECG classification algorithms fail to consider the predicting capacity, which refers to the capacity of triggering corresponding alarms before the occurrence of abnormalities. By comparing the morphology of signals which shows salient similarities to abnormal signals precedent to morbid heartbeats with annotated abnormal signal, the potential latent status between abnormal and normal signals is partially demonstrated. If abnormalities in cardiologist annotation are represented by red alarms, thus a system which is able to trigger a corresponding yellow alarm precedent to the abnormality can be considered as capable of predicting abnormalities as shown in Fig.1.2.

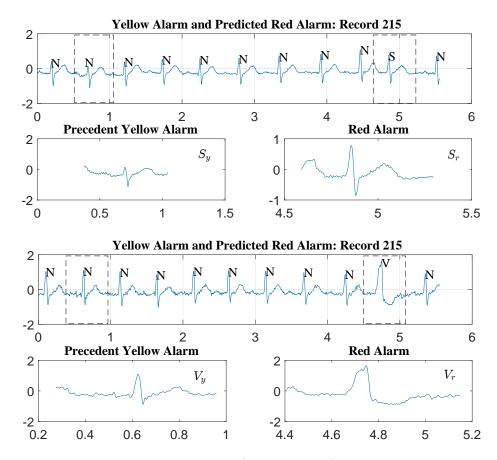


Figure 1.2: Potential latent abnormal status (yellow alarm) predict upcoming abnormality of same type.

Therefore, a method to quantify the level of signal similarity to abnormalities should be incorporated in ECG classification system.

1.4 Literature Review

Automatic analysis of ECG signals refers to the entire process from acquisition of signals to classification of samples. This process can be divided into five stages: ECG signal acquisition, preprocessing, fiducial peak detection and segmentation, feature extraction, predictive modeling (Fig. 1.3). There are researches focus on each single stage in the automatic analysis

system. Since the main objectives of this work are addressing problems in classification algorithms, the literature reviews in this section focus on studying existing methods proposed for stages before classification, conventional classification algorithms along with patient-specific classification systems.

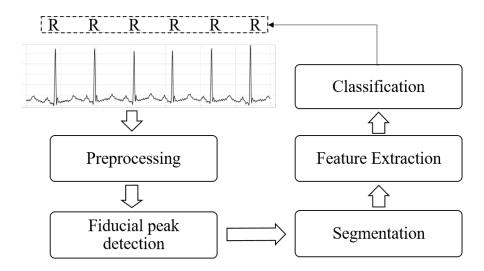


Figure 1.3: General structure of ECG classification system

1.4.1 ECG Signal Preprocessing

During data acquisition, ECG devices often introduce various noises including physiological noises (e.g. myoelectricity noise, breathe interference etc.) and non-physiological noises (e.g. power-frequency interference and electrode impedance interference) [12]. These noises often interfere with informative signals and thus influence ECG classification results. Therefore, ECG signal preprocessing mainly focus on the suppression of noise interference in ECG signals.

The ECG signal is in millivolt (mV) level with a frequency centered from 0 to 40 Hz [13]. Since ECG signal is comparatively weak with superposition of various noise, signal preprocessing is necessary step before classification. For this purpose, many researches have been

conducted targeting different noise component in ECG signal.

Generally speaking, ECG signal preprocessing methods includes finite impulse response (FIR) filtering, adaptive filtering, and modern signal processing filter methods such as wavelet transforms and neural networks. Ying -Wen Bai et al. compared different notch filters and concluded that equiripple notch filter performs the best on ECG signal. Lian et al. [14] designed a multiplier-free finite impulse response (FIR) filters to surpress biological and environmential noises with a low cost of power consumption. Sayadi et al. [15] proposed a modified extended Kalman filter with estimated hidden state variables to perform denoising and compression at the same time. Park et al. [?] designed a wavelet adaptive filter to reduce S-T segment distortion due to baseline drift and compared its performance with general adaptive filters. It is found that the wavelet adaptive filtering performance surpasses the general adaptive filters. In [16], the authors combined wavelet decomposition with Wiener filtering to filter noise by thresholding which is proved to outperform other thresholding denoising methods. Regarding various wavelet basis function, Singh et al. studied the optimal selection of basis function for ECG signal denoising [12]. By comparing the classification root mean square error using same classifier and different denoising methods, they concluded that Daubechies filter of order 8 performs the best for ECG classification system.

1.4.2 Fiducial Peak Detection and Segmentation

Fiducial Peak Detection and cardiac cycle segmentation are the basis for extracting important information from ECG signals since a ECG record is usually a continuous time series so the information about single cardiac cycle can only be obtained after segmenting the records. The accuracy and reliability of this stage directly determine the final performance of diagnosis and analysis.

Fiducial peak detection is also called ECG signal delineation, which aims at localize five characteristic peaks within one cardiac cycle. The most significant peaks are QRS complex consisting of Q, R and S peaks. The other two fiducial peaks are P wave before QRS complex and T wave after QRS complex. As shown in Fig.1.4, these five characteristic waves along with the onset and offset of QRS complex are often used the depict a cardiac cycle.

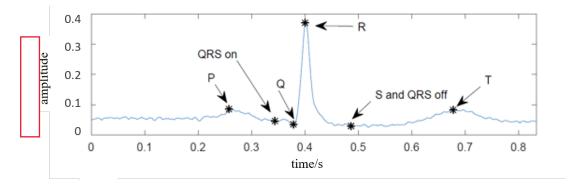


Figure 1.4: Fiducial peaks within one cardiac cycle

As QRS complex is the most prominent waves and it contains most of the information of ECG signal, most of the methods for ECG delineation detect QRS complex prior to the detection of other peaks. Afonso et al. [17] proposed a method using filter banks to detect QRS complex. In this method, signal is decomposed to several frequency band. Fiducial peaks are thus detected using its morphological features in the decomposed signals. Sadhukhan et al. [18] proposed a method of detecting R peak by comparing relative amplitude and double difference. The performance is validated on clinical ECG signal and proved to be promising. Some advanced machine learning techniques are also deployed to detect QRS complex. In [19], Support Vector Machine is used to train a predictive model for QRS complex detection and achived 99.93% accuracy. Other pattern recognition models such as Hidden Markov Models are investigated and proved to be effecient in modeling and detecting characteritic peaks in ECG signals [20]. Wavelet decomposition is also frequently adopted for signal delineation for the morphological similarity between wavelet basis function and QRS

complex. As the QRS complex power spectrum is centered at the range of 5 to 30 Hz, wavelet coefficients of corresponding scale levels are frequently used for delineation. In [21], QRS complexes are detected by thresholding wavelet coefficients at scales 1 to 4, then onset, offset and individual waves within QRS complexes are detected using morphological characters of coefficient, at scale 2. T and P waves are detected at scale 3 with similar method, Based on this method, some improvements have been proposed to eliminate false detection of R peaks by adding a fixed searching window of 160ms-in [22].

1.4.3 Feature Extraction and Classification

After localize the characteristic point within a cardiac cycle, the classification system needs to extract information in the signal so that the signal could be represented by a set of features. Since the objective of designing automatic classification system is to predict types of sample signal as precise as possible, the feature selection is usually performed to obtain a better performance and reduce the computation cost.

As the most significant wave within ECG signal, the information within QRS complexes are proved to be the most important features for a ECG classification system. Lagerholm et al. decompose QRS complexes with a set of Hermite basis function and the decomposition coefficient are deplyed as ECG features to train the Self-Organizing Map (SOM), which achieved an average error rate of 1.5% for 16 ECG types [23]. Prasad et al. [24] used discrete wavelet transform (DWT) to extract RR intervals between current beat and previous or next beats. The two RR-intervals serves as input for backpropagation neural networks training and the average accuracy of classifying 13 different arrhythmia types is 96.77%. De Chazal et al. proposed two set of features: morphology and heartbeat interval features. They used different combination of these features combined with Linear Discriminant Analysis to

classify ECG signal into five arrhythmia types and selected the optimal feature set according to classification performance. The result shows that the sensitivity of detecting two major arrhythmia types can be improved by feature selection. R. Ceylan *et al.* [25] included RR interval as the only ECG feature to train fuzzy clustering neural network and achieved 98.35% average detection rate for abnormal samples. Osowski *et al.* adopted High Order Statistics (HOS) and Hermite decomposition coefficients as feature set to classifity ECG signals with Support Vector Machine. Their final average error rate is at 1.82% [26].

1.4.4 Patient-Specific ECG Classification

The main drawbacks of the methods mentioned in the last section is the lack of interpatient model adjustment. In order to generalize the ECG classification system to clinical application, methods which are more robust to inter-patient signal variation are proposed to address this issue [8, 27–31].

Hu et al. proposed a patient-specific mixture of experts (MOE) classifier by incorporating personalized annotations provided cardiologists in the local classifier [27]. This methods achieves patient-adapting capacity but requires further input from human experts. This MOE approach achieved an accuracy of 94.0% for distinguishing ventricular beats from the other non ventricular types. Following the design of MOE, de Chazal and B. Reilly proposed an improved patient-adapting classifier by reducing the requirement of manual annotations to as few as 10 beats for training adaptive local classifier [28]. And Llamedo et al. designed an automatic classification system allowing but not depending on expert assistance [8]. By evolving a special block-based neural networks (BbNNs), Jiang et al. achieved accuracies of 98.1% and 96.6% for distinguishing ventricular ectopic beats and supraventricular ectopic beats from other types [29]. In [30], particle swarm optimization

(PSO) is combined with neural network to optimize network structure using patient specific training data. Based on 1-D convolutional neural networks (CNN), Kiranyaz et al. proposed a flexible algorithm which adjusts its parameters using information extracted from individual signal [31]. The classifier demonstrates consistent performance over different ECG records achieving an accuracy between 98% 99% for distinguishing VEBs from non-VEBs. (Acc = 98.9% Sen = 95.9% Spe = 99.4%). While this approach outperforms the aforementioned classification algorithms as it does not require expert further annotations, its performance reduces for some rare abnormal classes.

1.5 Contributions

A crucial drawback of these patient-specific classification systems in the literature are their deficiency of predicting abnormality in advance. Conventionally, classifiers associate each sample with a label and optimize performance according to the uniformity of proposed label and ground truth. While in many common applications this concept drives satisfying results, it does not meet the needs of SCD prediction.

One of the main objective of this work is to address the problem of forecasting by proposing two different feature space transformations to optimize the predicting capacity of personalized classifier for applications using biomedical signals. To specify this scenario, we assume that there are one normal status and multiple abnormal status for a signal while latent status exists for normalities with deviation towards abnormalities, indicating information regarding subsequent abnormalities. By distinguishing latent status, the designed automated system is capable to generate yellow alarms which indicates a high probability of proceeding presence of same type abnormalities. Therefore the contribution of this work can be summarized as:

- Proposed a novel self-adapting patient-adaptable framework which incorporate a personal classifier enabling predictive modeling
- Studied kernel-based method as spatial transformation with parameters optimized with Multi-Objective Particle Swarm Optimization for the purpose of deviation quantification
- Designed a controlled spatial transformation with deterministic mapping function to optimize cluster topology for predictive analysis
- Proposed a deviation quantification method based on cosine similarities which is able to generate predicting alarms for upcoming abnormalities

1.6 Organization of Thesis

In the following chapters, details of proposed classification frameworks are introduced along with the background of algorithms deployed in the framework. To start with chapter 2 provides a general introduction to the dataset used in this thesis and the general frameworks of the proposed classification system. Following this framework, chapter 3 further described the details of nonlinear transformation with kernel methods and presented the experimental results using kernel transformation. With the concept of nonlinear transformation, chapter 4 introduced an optimized spatial transformation method with a novel deterministic mapping function. The experimental result of the spatial transformation method is presented and studied in section 4.6. Finally, the experimental results of the methods proposed in chapter 4 and chapter 5 are compared. More importantly, the predicting capacity of systems are studied and analyzed. Based on the experimental results, we analyzed the potential future works in order to further optimize the system in both classification and predicting accuracy.

Chapter 2

Patient-Adaptable ECG Classification

Framework

2.1 Introduction

For decades, automatic ECG signal processing and analysis have been a controversial research topic. Studies carried out by scholars have proved that the development of automatic ECG analysis is conducive to the timely detection and therapeutic intervention of heart disease. However some major challenges need to be resolved before apply automatic ECG analysis as a highly reliable and fully automatic electrocardiogram processing system in clinical diagnosis. One of the most typical challenge is the inter-patient variation of ECG waveforms, which leads to inconsistent performance of ECG classification system. In this chapter, the basic framework of the proposed Patient-Adaptable ECG Classification will be discussed.

The goal of automatic ECG analysis is to determine the arrhythmia types for each signal sample. Continuous ECG signal is firstly segmented into individual segments which represent

heartbeat and processed by designed algorithms. The first sections in this chapter focus on the data preparation stage, which includes four steps: signal preprocessing, delineation and segmentation, and feature extraction. Following the data preparation Section 2.2 elaborate on the framework of a two staged hierarchical classifier. The classification system is patient adaptable by capturing the normal range for each individual. More specifically, in Section 2.3, the personal dynamic normal cluster method is discussed. One feature of this method is that the cluster can dynamically adapt to patient's ECG waveform change. In many application scenarios, the physicians need to monitor long-term real-time heart activity. The dynamic adjusting system is able to address the issue of intra-patient signal variation as well

2.2 ECG Data Preparation

2.2.1 Segmentation

Among a continuous ECG signal, the characteristics of a single cardiac cycle in the signal is generally considered as a sample in most machine learning applications. Therefore, it is necessary to segment ECG signal before feature extraction. Most of the methods in literature used wavelet analysis to detect the highest peak R wave in a cardiac cycle and then detect and segment it by the amplitude-frequency characteristics of other waves. In this study, the R feature peak segmentation method used in most of the literature is adopted. In Section ?? it was mentioned that a cardiac cycle consists of five basic characteristic peaks: P, Q, R, S, and T. Among them, the QRS complex is the most significant peak in one cycle. The energy of ECG signal in one cardiac cycle is mainly concentrated in the QRS complex. The QRS complex also contains important information that reflects the arrhythmia category [32]. Accurate detection of QRS complexes is of crucial importance for subsequent analysis. The

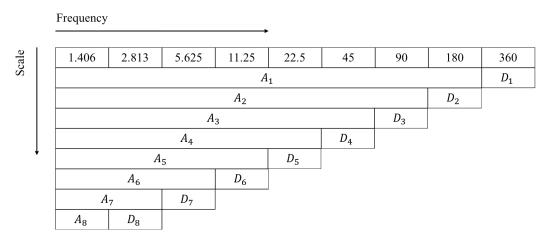


Figure 2.1: Frequency band of wavelet decomposition coefficient for MITDB signals

energy of the QRS complex is generally within the range of 5-25 Hz. Using this feature, wavelet analysis is applied to signals in MITDB with a sampling frequency of 360 Hz. With this sampling frequency, the decomposition coefficients and its frequency components are deduced in Fig.2.1 For ECG signals with a relatively low SNR, the QRS complex information can be extracted from the detail coefficients of level 5 (D_5) and level 6 (D_6)

The mother wavelet db4 is utilized at this stage due to its morphological similarity to QRS complexes. By superimposing D_5 and D_6 , the QRS complex information in the ECG signal can be characterized in a one-dimensional recombination signal $(QRS_DET = D_5 + D_6)$. Other fiducial peaks (P, QRS onset, Q, S, QRS offset and T waves for each cardiac circle) are localized according to the algorithm suggested by [32]. As shown in Fig.2.2

With the empirical values described in [32], we use 15% as the detection threshold. Since the width of most of the QRS complexes does not exceed 160ms, this work used a sliding window with a width of 160ms to detect the peaks in the QRS_DET . The window's step size was set to 200ms, given that the time interval between two adjacent heartbeat cycles does not exceed 200ms. Figure.2.3shows the waveform of QRS_DET and the corresponding window width of 160ms.

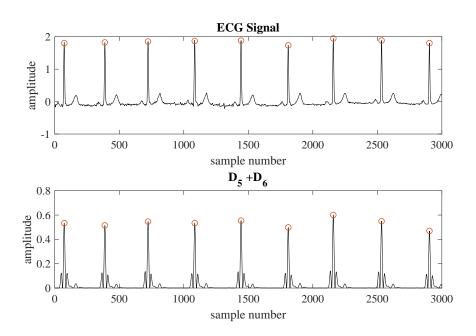


Figure 2.2: QRS detection with wavelet coefficients of level 5 and level 6

The false detection peaks are eliminated within a 160ms time window Fig.2.3

The T and P waves are outside the QRS window. By searching in a region from the start of a QRS to the end of its neighboring QRS, the P-wave is located in the highest positive peak in this region. In the same way, the position of the T wave is the position of the largest positive wave peak in the area from the end of the QRS to the next start of the next QRS. To sum up, the ECG signal in one heartbeat segment can be described by 7 feature points: P, QRS on, Q, R, S, QRS off and T.

Finally, using the location of the R wave, a segment of the ECG signal was divided into cardiac cycles. The starting position of a cardiac cycle is defined as the position 2/3 between the R wave of the cycle and the R wave of the previous cycle. The end position is defined as 2/3 of the R wave between the cycle R and the R wave of the following cycle position. The distance (RR) between each two adjacent R waves is h. The advantage of this method is that the computational complexity is low, and the scope of application is wide and it is

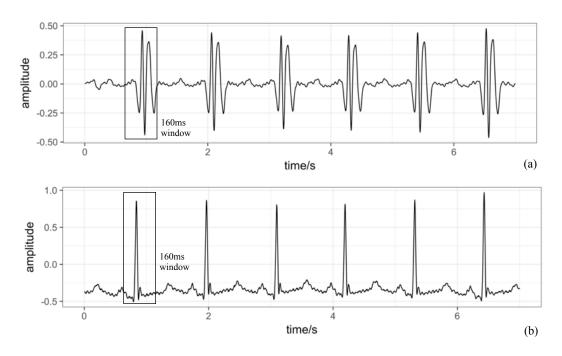


Figure 2.3: Window for detecting R peaks within QRS complexes

independent of individual patient differences.

However the quality of ECG signals provided by most of the portable ECG measuring instruments is very unstable. Signals transmitted through wireless communication systems will exhibit various unstable waveforms. Taking into account this feature, this paper performed subsequent feature extraction and analysis combined three consecutive ECG cycles, ie, one representative interval. In the subsequent analysis, an interval is a sample of data as shown in Fig.2.4.

For the purpose of training and evaluating classifier, MITDB is split into test (DS2) and training (DS1) set by balancing the four classes according to [33].

For each interval, a new annotation is generated by integrating all annotations of beats within the interval. Unless all beats are annotated as N within the interval, intervals will be annotated as the unique abnormality type within it. If two different abnormality types present within the same interval, this interval would be considered as transient and discarded.

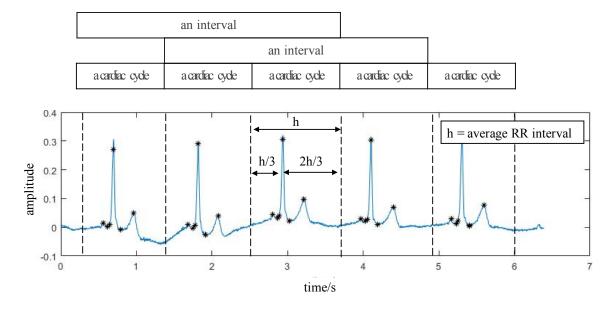


Figure 2.4: Interval and cardiac cycles

After segmentation and annotation generation, the total number of samples in training and test set is summarized in Table.2.1

2.3 Feature Extraction

The feature extraction of ECG signals plays a crucial role in the diagnosis of heart disease and has a great influence on the performance of subsequent automated classification systems. De Chazal et al. discussed in detail the classification results using waveform morphological features in [33]. As discussed in [34], the combination of three types of characteristics (ie, temporal, morphology, frequency domain) can distinguish between types of arrhythmias.

Table 2.1: Training and test datasets in MITDB.

	Number of intervals per AAMI class				
Evaluation Dataset	N	V	S	F	Total
DS1:Training	11721	2356	862	256	15195
DS2:Test	12633	2053	550	121	15357
Total	24354	4409	1412	377	30552

Table 2.2: Features extracted from ECG signal

Feature Type	SET1	SET2
Temporal Features	QRS duration, QT duration, PR duration	$\max(R_{i+1} - R_i),$ $\max(R_i - R_{avg})$
Morphological Features	max positive peak to second peak ratio	signal average energy, max pos- itive peak, max negative peak, peak to energy ratio
Frequency Domain Features	signal power level at 7.5Hz, 10Hz, 12.5Hz, 15Hz	

According to the related methods in literature, the different types of ECG signals mainly differ in the power level of the frequency band of 5 to 15 Hz in the segmented signal, and some other morphological features (such as the duration between the Q wave and the T wave, P The distance to the R wave, etc.) also shows a correlation with the signal class. Based on the interval segmentation method used in this paper, the feature extraction stage divided the three major types of features into periodic-based features (SET 1) and intervalbased features (SET 2). Here, SET1 includes the average and standard deviation of the corresponding features of the three cardiac cycles within a segment, and SET2 contains the overall characteristics of the time signal within the segment, so it is calculated once in intervals. Twenty-two feature quantities are extracted for each interval, which is then normalized to obtain a feature of zero-mean unit variance, and a 221 vector is used to represent the k-th interval. Summarized details are described in Table.2.2. The mean($R_{i+1} - R_i$) refers to the mean of the time interval between two adjacent R waves, while ($R_i - R_{avg}$) is the length of each cardiac cycle and the average cardiac cycle duration of the patient.

From Table. 2.2, it is explicit that these 22 feature are not completely independent of each

other. Therefore, in this work, Principal Component Analysis (PCA) is used to reduce the dimensions of features. Finally, 8 feature quantities were obtained after dimension reduction.

2.4 Classification Framework

In this section, a system is designed to address the classification and prediction tasks based on the processed ECG data. Based on our previous study and similar works which aimed to solve patient-specific classification problem, a two-staged structured which contains a global classifier and personalized classifier is retained as outline in this paper [8,27,28,35]. Moreover, the proposed algorithm incorporates a novel deviation quantification module depicted in details in the following section.

The framework of system is summarized by the flowchart in Figure 2.5. A pre-trained Global Classifier serves as preliminary classifier, it facilitates the following analysis in the system for it identifies samples with severe morbidity. Depending on applications, different conventional classification algorithms with low complexity can be considered as Global Classifier. Any abnormal labels generated by Global Classifier are considered as Red Alarms and do not require further processing.

However, as the objective of this study is to identify latent status between normality and abnormalities, the design focus on processing samples classified as N while deviating towards abnormal classes. For this purpose, a simple structured classifier is insufficient since sample numbers of N and other morbid classes are unbalanced in training set DS1 as shown in Table.2.1, which results in missed abnormalities. Thus a deviation detection module is added after the Global Classifier to specifically identify latent status using patient-specific normal cluster. In order to extract patient-specific information and adapt the classifier accordingly, the first 20% of total Normal samples of each patient are selected as initialization of Personal

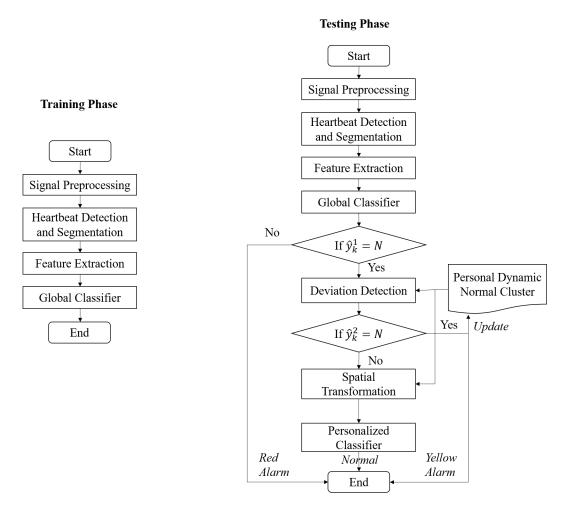


Figure 2.5: The general flowchart of proposed framework

Dynamic Normal Cluster \mathcal{N}_0^k . The binary classification of N versus non-N which includes all abnormal classes is completed by first calculating the following distance metrics:

$$R_i^{\text{max}} = \max_{\mathbf{x}_j \in \mathcal{N}_i^k, \mathbf{x}_k \in \mathcal{N}_i^k} \{ \sqrt{(\mathbf{x}_j - \mathbf{x}_k)^2} \}, \tag{2.1}$$

$$D_{\mathcal{X}}(\mathbf{x}_k(i)) = \underset{\mathbf{x} \in \mathcal{X}}{\operatorname{median}} \{ \sqrt{(\mathbf{x}_k(i) - \mathbf{x})^2} \},$$
(2.2)

$$D_{\mathcal{N}}^{\max}(\mathbf{x}_k(i)) = \max_{\mathbf{x} \in \mathcal{N}_i^k} \{ \sqrt{(\mathbf{x}_k(i) - \mathbf{x})^2} \},$$
 (2.3)

The following conditions are thus examined to verify if deviation of a sample is within the range defined by α . Considering the fact that some rare abnormalities are unlikely to be observed within the limit initialization time range, abnormal clusters: $\mathcal{S}, \mathcal{V}, \mathcal{F}$, which are composed of abnormal beats in DS1, are deployed as training data together with Personal Dynamic Normal Cluster \mathcal{N}_i^k in the following steps.

$$\begin{cases}
D_{\mathcal{N}}^{\max}(\mathbf{x}_{k}(i)) \leq \alpha R_{i}^{\max}, \\
D_{\mathcal{N}}(\mathbf{x}_{k}(i)) < D_{\mathcal{X}}(\mathbf{x}_{k}(i)) & \text{for } \mathcal{X} \in \{\mathcal{S}, \mathcal{V}, \mathcal{F}\}
\end{cases}$$
(2.4)

If a sample is confirmed as N in this module, it will be used to update the \mathcal{N}_i^k . Otherwise the system assumes that the sample have demonstrated deviation towards abnormal clusters and will pass it to the subsequent Personal Classifier with controlled nonlinear transformation. Using transformation with optimized parameters, Personal Classifier is able to discern the deviation to different morbid types regardless of the cluster topology within the original feature space. Details of the module will be presented in the next Section.

Overall, if given a sample x_k at time k, the proposed framework maps it to label $\hat{y}_k \in \{N, Y_V, Y_S, Y_F, R_V, R_S, R_F\}$, where N stands for normal, Y_V stands for ventricular yellow alarm, R_V stands for ventricular red alarm.

2.5 Personal Classifier

As the proposed system aims at predicting subsequent abnormalities by analyzing the deviation of sample signal, it's vital to quantify the deviations using topological characteristics of training data. For most of the ECG applications analysis is conducted within high-



Figure 2.6: The deviation analysis boundary restrict on latent status between normal and abnormal samples compared with the Global Classifier boundary

dimensional feature space, we therefore select Cosine Distance (defined in Eq.2.5) to quantify deviation.

$$d(\mathbf{v}, \mathbf{w}) = 1 - \frac{\mathbf{v}^T \mathbf{w}}{|\mathbf{v}||\mathbf{w}|} = 1 - \frac{\mathbf{v}^T \mathbf{w}}{\sqrt{(\mathbf{v}^T \mathbf{v})(\mathbf{w}^T \mathbf{w})}}$$
(2.5)

Consequently, relative deviations of sample from normal cluster(N) to other abnormal clusters (V, S, F) are defined by cosine distance between the vector $\mathbf{v}_k(i)$ (defined in Eq.2.6) to three vectors $\mathbf{v}_{\mathcal{X}}(i) = \mathbf{c}_{\mathcal{X}} - \mathbf{x}_k(i)$ where $\mathcal{X} \in \{\mathcal{S}, \mathcal{V}, \mathcal{F}\}$. In this case, smaller cosine distance represents higher alignments between normal cluster centroid $\mathbf{v}_k(i)$, current sample $\mathbf{x}_k(i)$ and the abnormal centroid $\mathbf{c}_{\mathcal{X}}$.

$$\mathbf{v}_k(i) = \mathbf{x}_k(i) - \mathbf{c}_N^k(i) = \mathbf{x}_k(i) - \sum_{\mathbf{x} \in \mathcal{N}_i^k} \mathbf{x} / |\mathcal{N}_i^k|, \tag{2.6}$$

Therefore the classification results of Personal Classifier $\hat{y}_k^2(i)$ is defined as follows:

$$\hat{y}_k^2(i) = \underset{\mathcal{X} \in \{\mathcal{S}, \mathcal{V}, \mathcal{F}\}}{\operatorname{argmin}} \{ d(\mathbf{v}_k(i), \mathbf{v}_{\mathcal{X}}(i)) \}$$
(2.7)

The measurement accuracy of cosine distance varies according to topology of clusters in feature space. Whereas the topology in feature space relies on feature extraction and feature selection. For example, as shown in Fig.2.7, in the original feature space overlaps and alignment of abnormal clusters may lead to inaccurate results of deviation quantification. In order to avoid the ambiguity, a topology where abnormal clusters encircle normal cluster is required.

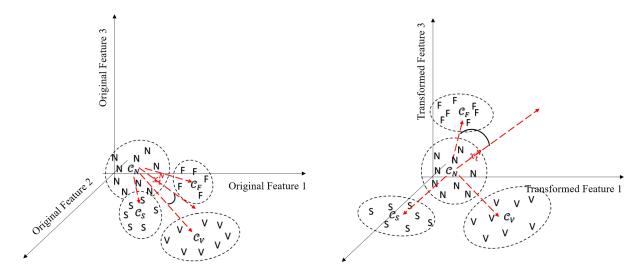


Figure 2.7: Left: illustration of cluster topology in original feature space; Right: illustration of cluster topology in feature space transformed with simple mapping function

For this purpose, two different spatial transformation methods are proposed in the next two chapters.

Chapter 3

Kernel-Based Nonlinear Spatial

Transformation

3.1 Introduction

As proposed in last chapter, objective of the system is to predict subsequent abnormalities by quantifying the deviation of sample. Whereas the original cluster topology in feature space Ω^d depends on the feature extraction stage g() and may lead to poor performance or even failure of predicting. An optimization using spatial transformation is addressed to solve this problem. More specifically, the designed method is able to maximize angles between vectors of normal cluster centroid to each abnormal cluster center centroids. In this chapter, a kernel based nonlinear spatial transformation is proposed to reshape the feature space to a symmetric topology which has the following features:

• Vectors pointing from normal centroid to different abnormal clusters centroids has no overlapping or cross with each other.

- Vectors pointing from normal centroid to different abnormal clusters centroids has maximum mutual cosine distance.
- The overlapping part of all clusters is minimized.

3.2 Kernel Method

First, given that clusters do not meet the symmetric features mentioned above, it's assumed in this chapter that new feature space obtained by non-linear reshaping is noted as $\Phi^{d'}$. This reshaping process is included in Personal Classifier stage (as shown in Fig.??) of the ECG classification system as described in chapter 2. The nonlinear mapping projects each sample \mathbf{x}_k onto a new vector \mathbf{z}_k in a higher dimension space, which can be considered as the union of subspace for each type: $\Phi^{d'} = \{\mathcal{N}, \mathcal{S}, \mathcal{V}, \mathcal{F}\}$, where d' > d

Kernel method has been widely applied in the machine learning algorithms, among which nonlinear support vector machine (SVM) has been deployed in numerous applications recently [36]. Nonlinear kernel method is most efficient when there is a nonlinear relationship between input and output variable. This nonlinear relationship assumption is usually true for ECG classification due to the complex feature vectors for ECG analysis. Therefore, it is practical to apply this method in the ECG analysis.

In nonlinear SVM, the algorithm minimize the expected error E[L(y, f(x))] so as to obtain classification function f, where L is the designated loss function, such as square error $(y - f(x))^2$ [37]. Based on input space $x_i \in \Omega$ and classifier mapping function f, we first define loss function through following formula:

$$\frac{1}{m} \sum_{j \in \mathbb{N}} L(y_j, f(x_j) + \gamma ||f||_K^2$$
 (3.1)

where, $||f||_K^2$ is the norm of f on H_K . If H_K is the Hilbert space of linear functions, f = w'x, $w \in \mathbb{R}^d$. Therefore the loss function can be written as:

$$\frac{1}{m} \sum_{j \in \mathbb{N}} L(y_j, w'x_j) + w'w \tag{3.2}$$

Positive constant γ known as regularization parameter, controls the balance between training error and the complexity (smoothness) of solution. SVM and other machine learning methods deploy kernel method with selected loss function [38]. In general, the solution to Eq.3.2 is as the following:

$$f(x) = \sum_{j \in \mathbb{N}} c_j K(x_j, x) \tag{3.3}$$

where c_j is a real number, K is kernel function, such as polynomial function: $K(x,t) = (x't)^r$, $x, t \in \mathbb{R}^d$

Hibert space H_K is usually defined base on feature mapping concept, $\Psi: \Phi^d \to \Phi^{d'}$ where $\Phi^{d'}$ is Hilbert space and $<,>_{\Phi}$ represents inner product. Therefore, Eq.3.3 can be also written as:

$$f(x) = \sum_{j \in \mathbb{N}} c_j \Psi(x_j, x) \tag{3.4}$$

For nonlinear kernel function method, the selection of kernel function plays a decisive role. An effective kernel function generally needs to satisfy Mercer theorem [39]. This means the matrix defined by function $\Psi(x_j, x)$ is symmetric positive semi-definite. In general, the selection of kernel function depends on all observations in input space. However, in effect, kernel functions with simple expression, such as polynomial kernel function, Gaussian kernel function and exponential kernel function are usually preferred than complicated kernel function for its simplicity and consistency. Polynomial kernel function is usually applied on normalized data for its explicit expression and steady performance. Whereas degree of freedom of polynomial kernel function is comparatively high which leads to a large of parameters. Gaussian kernel function, a very classic robust radial function, has high robustness when the data includes strong noise. Nevertheless, since Gaussian kernel actually projects samples to a infinite dimensional space, it's difficult to visualize projected observations and interpret the result. Moreover its performance is greatly affected by parameter selection.

In this work, the representative polynomial kernel is selected for the purpose of validating the method and explaining the effect of nonlinear kernel method and optimization on the feature space reshaping. The kernel functions can be written in the following format:

$$\mathbf{z}_{k} = \boldsymbol{\Psi}_{\mathbf{w}}(\mathbf{x}_{k}) = \begin{bmatrix} w_{1} \\ w_{2} \\ \vdots \\ w_{d'} \end{bmatrix} \circ \begin{bmatrix} \psi_{1}(\mathbf{x}_{k}) \\ \psi_{2}(\mathbf{x}_{k}) \\ \vdots \\ \psi_{d'}(\mathbf{x}_{k}) \end{bmatrix}$$

$$(3.5)$$

The example above shows that regardless the selection of kernel function, parameter optimization will play a critical role. For instance, in this work, the process of spatial topology optimization is accomplished by adjusting the coefficients of fixed polynomial basis functions $\psi()$. Since the number of parameters to adjust is very large when the order of polynomial functions is high, exhaustive algorithms are not practical for parameter optimization. Therefore, it's necessary to implement a heuristic optimization algorithm, in which parameters are obtained by maximizing or minimizing a loss function. More specifically, the nonlinear reshaping in this work aims to adjust mapping coefficient $\mathbf{w} = [w_1, w_2, \dots w_d]^T$ to achieve the ideal symmetry of clusters in the reshaped feature space while maintaining the distance between clusters.

3.3 Multiobjective Optimization

3.3.1 Objective Functions

To illustrate the optimization problem, if we assume the original feature space is the 2-dimension space Ω^2 , then the mapping kernel function may adopt second-order polynomial function as follows:

$$\mathbf{x} = [x_1 \ x_2]^T, \quad \mathbf{w} = [w_1 \ w_2 \ \dots \ w_5]^T, \quad d = 2, \quad d' = 5$$

$$\psi_1(\mathbf{x}) = x_1, \psi_2(\mathbf{x}) = x_2, \psi_3(\mathbf{x}) = x_1^2, \psi_4(\mathbf{x}) = x_2^2, \psi_5(\mathbf{x}) = x_1 x_2$$
(3.6)

Similar to the loss function in the standard nonlinear kernel method, the following objective functions can be used to represent the symmetrical structure to be obtained:

$$o_{1}(\mathbf{w}) = \frac{1}{\min_{c,d=2,...,p \text{ and } c\neq d}} \{d(\mathbf{v}_{\mathcal{X}_{c}}, \mathbf{v}_{\mathcal{X}_{d}})\}$$

$$o_{2}(\mathbf{w}) = \frac{SW}{SB} = \frac{\sum_{c=1}^{C} \sum_{\mathbf{z} \in \mathcal{X}_{c}} (\mathbf{z} - \mathbf{c}_{\mathcal{X}_{c}})^{T} (\mathbf{z} - \mathbf{c}_{\mathcal{X}_{c}})}{\sum_{c=1}^{C} \sum_{d=1, d\neq c}^{C} (\mathbf{c}_{\mathcal{X}_{c}} - \mathbf{c}_{\mathcal{X}_{d}})^{T} (\mathbf{c}_{\mathcal{X}_{c}} - \mathbf{c}_{\mathcal{X}_{d}})}$$

$$(3.7)$$

The maximization of pair cosine distance between vectors $\mathbf{v}_{\mathcal{X}_c}$ connecting the centroid of the normal class to the abnormal classes \mathcal{X}_c can be gained by minimizing $o_1(\mathbf{w})$. In the formula, vectors used to measure symmetry is defined by abnormal sample sets in training set DS1 and the personal normal cluster as follows:

$$\mathbf{v}_{\mathcal{X}_i} = \mathbf{c}_N^k - \mathbf{c}_{\mathcal{X}_i} \tag{3.8}$$

On the other hand, $o_2(\mathbf{w})$ represents the ratio of within cluster variance to between-cluster variance, hence controls the separation between clusters. Cosine distance is defined by Eq.2.5 and these objective functions are deduced from discrimination function of personal classifier in Eq.2.7. By minimizing these two objective functions at the same time, the algorithm

eliminates the ambiguity while applying cosine distance and hence improve the predicting capacity.

3.3.2 Multiobjective Particle Swarm Optimization

It should be noted that $o_1(\mathbf{w})$ and $o_2(\mathbf{w})$ are not necessarily independent to each other. Thus the optimization problem here means the jointly minimizing of $o_1(\mathbf{w})$ and $o_2(\mathbf{w})$ subject to constraint condition $|w|_2 = 1$. Since this is a non-convex multiobjective optimization problem, closed form solution or optimization methods for convex function are not suitable for this problems. Therefore, this paper adopts multiobjective particle swarm optimization (MOPSO) algorithm to solve it.

Particle Swarm Optimization (PSO) has the advantage of fast-converging, heuristic searching and easy implementation [40,41]. Therefore, researchers have already started investigating in extending PSO to multiobjective optimization problems. In the framework of MOPSO, the goal is to solve the typical Pareto optimization problem with the structure of PSO. In other word, it aims at solving optimization problem with two or more conflicting objective functions by approximating the Pareto front.

Among all MOPSO algorithms in literature, the algorithm proposed by Coello Coello and Lechug facilitates the implementation and improved the performance compared to other methods [40]. Therefore, this algorithm is adopted in this work. One featured design of this algorithm is the external repository in which all Pareto optimized particles in each swarm is recorded for each iteration. The configuration of repository members are stored and used as an optimal approximation of the Pareto front of the problem as they converge to the actual Pareto front as proved in [40]. Fig.3.1 generated by optimizing $o_1(\mathbf{w})$ and $o_2(\mathbf{w})$ demonstrates the repository members are Pareto optimal than other particles in the

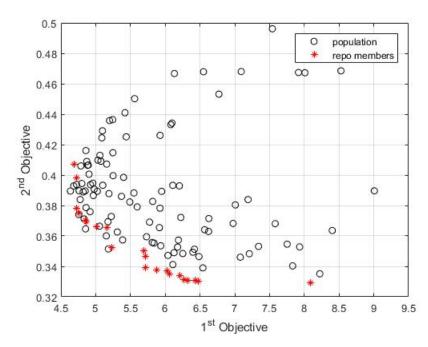


Figure 3.1: Particles stored in external repository approximate the Pareto front

objective function space and they converge to the Pareto front.

With the concept of Pareto front, we further demonstrate the impact of applying kernel functions in this multiobjective optimization problem by comparing Pareto front before transformation with kernel method and the Pareto front with nonlinear reshaping.

As shown in Fig.3.2, the Pareto front in feature space after nonlinear reshaping is superior obviously to that of Pareto front with linear combination of original data. The kernel function used in this comparison is a third-order polynomial kernel function as formulated in Eq.3.6. The result shows that nonlinear kernel method possess a higher degree of freedom to in multiobjective optimization. In other words, kernel method combined with multiobjective particle swarm optimization algorithm can improve the spatial topology of clusters.

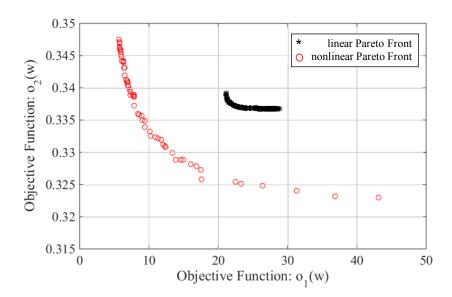


Figure 3.2: Increase of degree of freedom in optimization is proved by comparing the Pareto fronts generated by linear and nonlinear basis function

3.4 Experimental Results

This section will introduce the experimental result generated with test set DS2 of MITDB to show the performance of the proposed method in this Chapter. Here, we firstly map the original 22-dimensional feature vectors representative of cardiac segments into a 8-dimensional vectors $\mathbf{x}_{8\times 1}$ using principal component analysis (PCA). In order to realize the non-linear transformation in (3.5), we use polynomial function of order 3. Taking the computational cost into account and also to avoid over-fitting of higher-order data samples, we only retain 32 terms (8 second order pure terms x_i^2 , 8 third order pure terms x_i^3 , 8 second order cross terms $x_i x_j$, and 8 pure third order cross terms $x_i x_j^2$) and randomly discard the rest of cross terms. Therefore, the mapped vectors $\mathbf{z}_{32\times 1}$ include a total of 32 terms as follows.

$$\mathbf{z} = \{x_i^2 | i = 1, 2 \dots 8\} \cup \{x_i^3 | i = 1, 2 \dots 8\} \cup \{x_i x_j | i, j = 1, 2 \dots 8, i \neq j\} \cup \{x_i^2 x_j | i, j = 1, 2 \dots 8, i \neq j\}$$

$$(3.9)$$

The performance of the proposed system in this paper is tested with DS2 excluding record 232, for this record has only 7 normal samples $y_k = N$. In total, 21 records are tested.

Table 3.1, shows the performance of the proposed method in classification of ECG signal segments. In order to average the results over all recordings, we present the median, interquartile range (IQR), mean and standard deviation of accuracy (AC), sensitivity (SE) and specificity (SP). The results are promising and the median of accuracy for all classes are in the range of 88% - 99%. Sensitivity and specificity of the proposed method exhibits similar ranges. The mean accuracy is at least 86% excluding class V. Therefore, this system is very less likely to miss an important alarm or to report false alarms.

Table 3.1: Classification results of the proposed method.

Class N	median(%)	IQR(%)	mean(%)	std (%)
AC	94.8	19.52	86.62	18.55
SE	97.21	17.36	87.47	19.26
class V	median(%)	IQR(%)	mean(%)	std (%)
AC	86.11	27.54	76.41	22.81
SP	99.71	11.22	90.18	18.52
class S	median(%)	IQR(%)	mean(%)	std (%)
AC	99.28	2.24	98.29	2.57
SP	99.64	22.17	97.56	6.06
class F	median(%)	IQR(%)	mean(%)	std (%)
AC	97.91	8.2	93.85	7.84
SP	100.00	0.03	99.12	3.6

More importantly, the predicting ability of the proposed method is worthy of evaluating separately. In order to quantify the conditional probability of observing a red alarm after a

Table 3.2: Predictive power of yellow alarms: A yellow alarm increases the chance of observing a red alarm of the same type.

		ıber o ormalit	of next			ability abno	of of ormality	
secondary abnor- malities	V_y	S_y	F_y	Tota	$1 V_y$	S_y	F_y	Total
True V	38	23	35	96	75	75	61	67
True S	11	10	8	29	21	29	14	20
True F	2	2	14	18	4	6	25	13

preceding yellow alarm of similar type in (??), we count the number of predicted samples as follows:

$$P(\hat{y}_{k+i} = X_r | \hat{y}_k = X_y) = \frac{\text{\# of } y_{k+i} = X \text{ after } \hat{y}_k = X_y}{\text{\# of true alarms after } \hat{y}_k = X_y}$$

$$P(\hat{y}_{k+i} = X_r) = \frac{\text{\# of true alarm of type } X (y_k = X)}{\text{\# of all true alarms}}$$
(3.10)

The summary of results for all 21 test records is presented in Table. 4.4. The last 4 columns of the Table. 4.4 show the probability of having a subsequent true abnormality of any type after observing a yellow alarm. These results confirm the predictive power of yellow alarms. For instance, the absolute probability of observing a segment with abnormal classes V, S, and F is respectively $\frac{96}{96+29+18}=67\%$, $\frac{29}{96+29+18}=20\%$ and $\frac{18}{96+29+18}=13\%$, based on their relative frequencies. However, these probabilities after observing a yellow alarm of type Vp are respectively $\frac{38}{38+11+2}=75\%$, $\frac{11}{38+11+2}=21\%$ and $\frac{2}{38+11+2}=4\%$. This means that the probability of observing a red alarm of type V is 75%-67%=8% higher than its absolute probability. The same trend holds for other yellow alarms as well. The results suggest a more in-depth study of the concept of yellow alarms for heart monitoring.

3.5 Conclusions

This chapter explains the nonlinear reshaping with kernel function and multipurpose particle swarm optimization method. With the kernel method adopted in SVM, we utilize a group of nonlinear kernels to reshape the input feature space and map it to high-dimensional feature space, which meets two conditions, namely, maximum separation between cluster and maximum cosine similarities between abnormal clusters.

In this chapter we adopt multipurpose particle swarm optimization method to optimize parameters of kernel function. Result shows that Pareto front produced by kernel method in the objective function space is apparently optimal to that produced by linear combination of features. The results verify the accuracy of the proposed method with a classification accuracy range of 88% - 99% for different ECG records in publicly available MIT-BIH database.

Above all the proposed methodology demonstrates a potential to add detailed information about sample deviation upon conventional machine learning algorithm. We tested this system with ECG signal and observed promising results, but this method is not bound to this application.

Chapter 4

Controlled Spatial Transformation With Deterministic Mapping Function

4.1 Introduction

In Chapter 2, we elaborated on the details of patient adaptable ECG classification framework. In Chapter 3, the specific spatial topology of normal and abnormal clusters was analyzed in order to model the trajectory of ECG samples evolving from normal status to abnormal. These two methods share a common concept of modeling the intermediate states from annotated normal to abnormal state.

While the aforementioned systems demonstrated capacity of predicting upcoming abnormalities, it's challenging to interpret the mechanisms of the systems and thus hindering the generalization of predictive warning to other applications of biomedical signals. Therefore, the main object of this chapter is developing a classification system with abnormality predicting capacity based on spatial topology studied in [35].

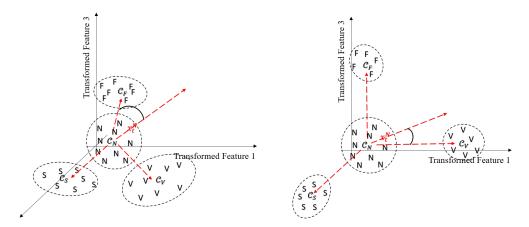


Figure 4.1: Left: illustration of cluster topology in feature space transformed with simple mapping function; Right: illustration of cluster topology in feature space transformed with optimized mapping function

Having analyzed the symmetric encircled topology in the previous chapter, we proposed a novel spatial transformation based predictive modeling system to assist cardiologist take advanced therapeutic Interventions. In this method, we further optimize the topology and spatial geometry of clusters in the feature space by reducing the within cluster variance after spatial transformation. As shown in 4.1, through this improvement, the predicting accuracy of personal classifier can be further improved.

4.2 Hyper-Spherical Coordinates

In the precedent chapter, the spatial transformation module was implemented with kernel function and heuristic optimization algorithm. The system performance was proved to be promising. However it's difficult to select the appropriate kernel as spatial mapping function due to the high variety of kernel functions and the limitation of dimensionality.

In order to address this problem, a novel deterministic spatial mapping function is proposed based on hyper-spherical coordinates in this work. Since hyper-spherical coordinates consists of angles and radius, this information together with the cluster topology are used to determine the mapping function parameters.

Hyper-spherical coordinates (n-dimensional spherical coordinates) and its mapping to Cartesian coordinates are first elaborated in [42]. If \mathbf{x} is a sample vector in n-dimensional feature space, where its Cartesian coordinates is $(\xi_1, \, \xi_2 \, ... \xi_n)$, then its corresponding hyper-spherical coordinates can be obtained through Eq.4.1 which is originally derived through its reverse mapping (Eq.4.2) using $\sin(\arccos(x)) = \sqrt{1-x^2}$

$$r = \sqrt{\xi_n^2 + \xi_{n-1}^2 + \dots + \xi_2^2 + \xi_1^2}$$

$$\theta_1 = \arccos \frac{\xi_1}{\sqrt{\xi_n^2 + \xi_{n-1}^2 + \dots + \xi_1^2}}$$

$$\theta_2 = \arccos \frac{\xi_2}{\sqrt{\xi_n^2 + \xi_{n-1}^2 + \dots + \xi_2^2}}$$

$$\vdots$$

$$\theta_{n-2} = \arccos \frac{\xi_{n-2}}{\sqrt{\xi_n^2 + \xi_{n-1}^2 + \xi_{n-2}^2}}$$

$$\theta_{n-1} = \begin{cases} \arccos \frac{\xi_{n-1}}{\sqrt{\xi_n^2 + \xi_{n-1}^2}} & \xi_n \ge 0 \\ -\arccos \frac{\xi_{n-1}}{\sqrt{\xi_n^2 + \xi_{n-1}^2}} & \xi_n < 0 \end{cases}$$

$$\xi_{1} = r \cos(\theta_{1})$$

$$\xi_{2} = r \sin(\theta_{1}) \cos(\theta_{2})$$

$$\xi_{3} = r \sin(\theta_{1}) \sin(\theta_{2}) \cos(\theta_{3})$$

$$\vdots$$

$$\xi_{n-1} = r \sin(\theta_{1}) \cdots \sin(\theta_{n-2}) \cos(\theta_{n-1})$$

$$\xi_{n} = r \sin(\theta_{1}) \cdots \sin(\theta_{n-2}) \sin(\theta_{n-1}).$$

$$(4.2)$$

where
$$0 \le \theta_j \le \pi$$
, $j = 1, ..., n - 2$; $0 \le \theta_{n-1} \le 2\pi$; $0 \le r < \infty$

4.3 Orthogonalization

To simplify algorithms, in this paper the topology of clusters in feature space is approximated by relative topology of cluster centroids: $\mathbf{c}_N^k, \mathbf{c}_{\mathcal{V}}, \mathbf{c}_{\mathcal{S}}, \mathbf{c}_{\mathcal{F}}$. Furthermore, as we assume that samples deviates from normality to abnormality and spatial topology stays unchanged if simple translation is applied to data, the cluster topology in original feature space can be simply represented by the following matrix:

$$C = \begin{bmatrix} \mathbf{c}_{\mathcal{V}} - \mathbf{c}_{N}^{k} \\ \mathbf{c}_{\mathcal{S}} - \mathbf{c}_{N}^{k} \\ \mathbf{c}_{\mathcal{F}} - \mathbf{c}_{N}^{k} \end{bmatrix} = \begin{bmatrix} \mathbf{V}_{\mathcal{V}\mathcal{N}} \\ \mathbf{V}_{\mathcal{S}\mathcal{N}} \\ \mathbf{V}_{\mathcal{F}\mathcal{N}} \end{bmatrix}$$

$$(4.3)$$

As shown in Fig.4.1, in order to improve Personal Classifier and avoid ambiguity, a topology with maximum separation between three vectors in \mathcal{C} is ideal. In order to lower computing

complexity in high dimension, the algorithm aims at transforming vectors in \mathcal{C} to orthogonal vectors with deterministic functions. Therefore, the method proposed in [43] based on the well known orthogonalization method Gram-Schmidt in [44] is deployed. Hence, in the first step of the function a vector of the centroids is fed to the orthogonalization process as follows:

$$C^{\perp} = \operatorname{Gram-Schmidt}(C) = \begin{bmatrix} \mathbf{V}_{\mathcal{V}\mathcal{N}}^{\perp} \\ \mathbf{V}_{\mathcal{S}\mathcal{N}}^{\perp} \\ \mathbf{V}_{\mathcal{F}\mathcal{N}}^{\perp} \end{bmatrix}$$
(4.4)

where \mathcal{C}^{\perp} is the matrix of Cartesian coordinates orthogonalized vectors. The hyper-spherical coordinates \mathcal{C}^{\perp}_* of same orthogonalized vectors, which reveals the angular change from original vectors to the ideal orthogonal vectors, are calculated subsequently using Eq.4.1.

$$\mathcal{C}_{*}^{\perp} = \begin{bmatrix} r_{\mathcal{V}\mathcal{N}}^{\perp} & \theta_{1_{\mathcal{V}\mathcal{N}}}^{\perp} & \dots & \theta_{n-1_{\mathcal{V}\mathcal{N}}}^{\perp} \\ r_{\mathcal{S}\mathcal{N}}^{\perp} & \theta_{1_{\mathcal{S}\mathcal{N}}}^{\perp} & \dots & \theta_{n-1_{\mathcal{S}\mathcal{N}}}^{\perp} \\ r_{\mathcal{F}\mathcal{N}}^{\perp} & \theta_{1_{\mathcal{F}\mathcal{N}}}^{\perp} & \dots & \theta_{n-1_{\mathcal{F}\mathcal{N}}}^{\perp} \end{bmatrix}$$
(4.5)

4.4 Spatial Mapping Function

After obtaining the original spherical coordinates $(\mathbf{V}_{\mathcal{V}\mathcal{N}}, \mathbf{V}_{\mathcal{S}\mathcal{N}}, \mathbf{V}_{\mathcal{F}\mathcal{N}})$ and orthogonalized spherical coordinates $(\mathbf{V}_{\mathcal{V}\mathcal{N}}^{\perp}, \mathbf{V}_{\mathcal{S}\mathcal{N}}^{\perp}, \mathbf{V}_{\mathcal{F}\mathcal{N}}^{\perp})$, the goal is to design a mapping function $\mathbf{F}: \mathbf{R}^n \to \mathbf{R}^n$ from original coordinates to the orthogonal ones which is equivalent to the ideal cluster topology.

In Gram-Schmidt algorithm, the very first vector in the input vector array serves as a reference vector and remains unchanged in the orthogonal vector array. As a result, $\mathbf{V}_{\mathcal{V}\mathcal{N}} =$

 $\mathbf{V}_{\mathcal{V}\mathcal{N}}^{\perp}$ and equivalently \mathbf{F} can be defined by the following equations:

$$\mathbf{F}(\mathbf{V}_{SN} - \mathbf{V}_{VN}) = \mathbf{V}_{SN}^{\perp} - \mathbf{V}_{VN}^{\perp} = \mathbf{V}_{SN}^{\perp} - \mathbf{V}_{VN}$$

$$\mathbf{F}(\mathbf{V}_{FN} - \mathbf{V}_{VN}) = \mathbf{V}_{FN}^{\perp} - \mathbf{V}_{VN}^{\perp} = \mathbf{V}_{FN}^{\perp} - \mathbf{V}_{VN}$$

$$(4.6)$$

Furthermore, since orthogonality is Independent to radius r, \mathbf{F} only needs to apply to n-1 dimensions which includes all angles $(\theta_1,\ldots,\theta_{n-1})$ and coordinate r remains unchanged before and after mapping. Consequently, determination of \mathbf{F} can be decomposed to the determination of n-1 functions: $f_i: \mathbf{R} \to \mathbf{R}, i = 1 \dots n-1$ with constraints in Eq.4.6 and boundary constraints. Note the $\mathbf{V}_{SN} - \mathbf{V}_{VN}$ as Δ_{SV} and its ith angular dimension of as δ_{isv} . Same notation is applied on $\mathbf{V}_{FN} - \mathbf{V}_{VN}$. Hence for each angular dimension i, f_i is determined by $(\delta_{isv}, \delta_{isv}^{\perp})$ and $(\delta_{irv}, \delta_{irv}^{\perp})$, together with two boundary constraints In order to maintain the simplicity and linearity of the mapping function, f_i needs to be continuous and monotonic. For this purpose, periodicity of angular dimension is used to determine the boundary constrains and the problem is transformed into a curve fitting one. For example, if linking $(\delta_{isv}, \delta_{isv}^{\perp})$ and $(\delta_{irv}, \delta_{irv}^{\perp})$ results in a monotonically decreasing line, the boundary constrains would be $(\pi, 0)$ and $(0, \pi)$. Conversely, if it results in a monotonically increasing line, the boundary constrains would be (0, 0) and (π, π) . Same rules applies on the last angular dimension where period is 2π instead of pi.

The simplest candidate function for f_i , which connect two boundary points and two target points in the 2-D plane would be a linear spline as shown in Fig.4.2

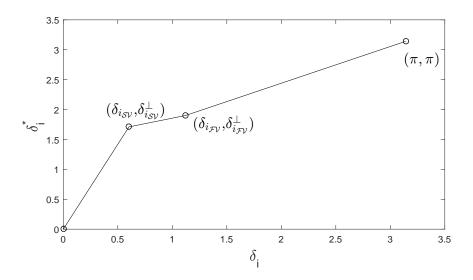


Figure 4.2: The simple mapping function

4.5 Optimized Mapping Function

The mapping function in Fig.4.2 demonstrated monotonicity and continuity, which is a part of targeted characteristics as an ideal mapping function for f_i . However there exists some drawbacks in the simple mapping function. First of all, the function is not differentiable at two target points $(\delta_{i_{SV}}, \delta_{i_{SV}}^{\perp})$ and $(\delta_{i_{FV}}, \delta_{i_{FV}}^{\perp})$, which will lead to cluster deformation. Secondly, since the mapping function is applied on angular dimension, linearity of each spline in the function will bring on non-convex clusters in Cartesian coordinates after mapping. In order to avoid deformations and preserve within cluster geometry, it is required to have a region for each cluster centroids within which the function maps the data to a equal or even smaller range in Cartesian coordinates. In the other word, the samples close to the centroids has to be close to the same corresponding centroids in the mapped space. In order to avoid deformations, an optimized mapping function is proposed.

To compromise three constrains, a function which satisfies the following mathematical conditions is studies:

- mapping function derivatives around centroids 0 and $\delta_{i_{SV}}$ and $\delta_{i_{FV}}$ are small
- mapping function derivatives between two centroids large
- quasi differentiable at all points

Therefore, we proposed the basis function p satisfies the aforementioned constrains. The function p is composed of two parts: h(x) and g(x). The boundaries that each function activates are defined as target points $(\delta_{i_{\mathcal{S}\mathcal{V}}}, \delta_{i_{\mathcal{S}\mathcal{V}}}^{\perp})$ and $(\delta_{i_{\mathcal{F}\mathcal{V}}}, \delta_{i_{\mathcal{F}\mathcal{V}}}^{\perp})$ along with the mid-points between target points. The lower boundary mid points are noted as (γ, γ^{\perp}) . The upper boundary mid points are noted as $(\epsilon, \epsilon^{\perp})$. To ensure the continuity of mapping function f, we have

$$(\epsilon_{i_{\mathcal{S}\mathcal{V}}}, \epsilon_{i_{\mathcal{S}\mathcal{V}}}^{\perp}) = (\gamma_{i_{\mathcal{F}\mathcal{V}}}, \gamma_{i_{\mathcal{F}\mathcal{V}}}^{\perp}) = (\frac{\delta_{i_{\mathcal{S}\mathcal{V}}} + \delta_{i_{\mathcal{F}\mathcal{V}}}}{2}, \frac{\delta_{i_{\mathcal{S}\mathcal{V}}}^{\perp} + \delta_{i_{\mathcal{F}\mathcal{V}}}^{\perp}}{2})$$
(4.7)

Therefore, the two piecewise functions h(x) and g(x) are defined as follows:

$$K_h = \frac{\epsilon^{\perp} - \delta^{\perp}}{e^{\alpha(\epsilon - \delta, 0)^{+}} - 1}$$

$$h(x) = K_h[e^{\alpha(x - \delta, 0)^{+}} - 1] + \delta^{\perp}$$

$$(4.8)$$

$$K_{g} = \frac{\gamma^{\perp} - \delta^{\perp}}{e^{\alpha(-\gamma + \delta, 0)^{+}} - 1}$$

$$g(x) = K_{g}[e^{\alpha(\delta - x, 0)^{+}} - 1] + \delta^{\perp}$$
(4.9)

Applying p on two target points $(\delta_{i_{SV}}, \delta_{i_{SV}}^{\perp})$ and $(\delta_{i_{FV}}, \delta_{i_{FV}}^{\perp})$ will result in a smooth curve as

shown in Fig.4.3

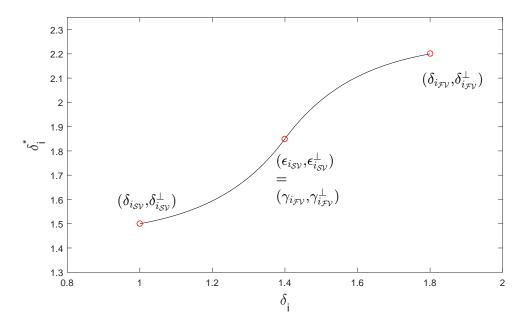


Figure 4.3: Optimized Piecewise Interpolate Function p

And applying the piecewise interpolate function p at all four points: $(\delta_{i_{SV}}, \delta_{i_{SV}}^{\perp})$, $(\delta_{i_{FV}}, \delta_{i_{FV}}^{\perp})$ and two boundary points, the final result is shown in Fig.4.4

4.6 Experimental Results

In this section, the performance of proposed method is evaluated according to two aspects. We will first analyze the classification performance of the system and demonstrate the comparison with other representative ECG classifiers. Furthermore, the classification results are partitioned into two sets: red alarms generated by Global Classifier and final results by combining yellow and red alarms. Impacts of Personal Classifier on final results will be studied. Finally the prediction performance which is typical in the proposed system is evaluated.

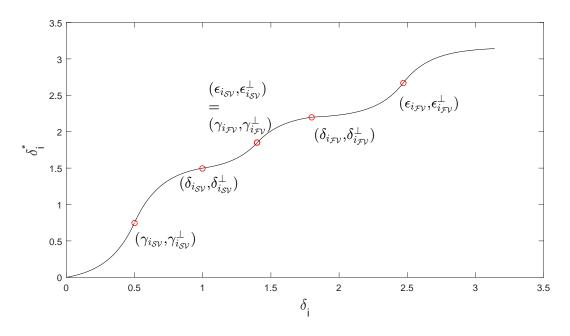


Figure 4.4: Optimized Mapping Function f

4.6.1 Classification Performance

The experimental results are evaluated with the classification performance of 4 AAMI ECG classes using the test subset of MITBIH Arrhythmia DS2. Originally DS2 contains 15357 samples after feature extraction. While training Personal Classifier, the first 20% of total normal samples serve as initialization samples for Personal Dynamic Normal Cluster. Therefore, all samples before the last initialization normal sample should be excluded from test set for each records. Consequently the actual test set contains 12414 samples in total consisting of 10105 type-N, 1702 type-V, 508 type-S and 99 type-F samples.

To present the result, we select weighted k-Nearest Neighbors where k = 10 as Global Classifier as it's comparatively simple and representative among low complexity models. Parameter α in the deviation detection module is set to 1 for test purpose.

Table 4.1 summarized the cumulated confusion matrix for all records in the test set. In order to compare the result of Global Classifier and combined result, the sample numbers

Table 4.1: Cumulated Confusion Matrix for All Records in DS2

		Ground Truth											
		N	V	S	F								
	N	9255(10076)	21(38)	72(90)	1(5)								
Result	V	657(22)	1678(1663)	8(2)	9(7)								
	S	71(6)	3(1)	417(416)	0(0)								
	F	122(1)	0(0)	11(0)	89(87)								

Table 4.2: Classification Performance and Within-Set Variation of Proposed System

statistics	N			V			S			F		
Statistics	Ac	Se	Sp									
cumulated	92.4	91.59	95.93	94.38	98.59	93.71	98.67	82.09	99.38	98.85	89.9	98.92
median	94.45	92.21	95.42	96.17	99.55	95.71	99.38	80.65	99.84	99.11	90.91	99.11
IQR	6.33	10.08	11.91	5.17	1.64	8.62	1.76	19.35	0.61	1.58	23.33	1.49

are presented in the following format: combined(globaled). In order to measure classification performance, we adopted four metrics proposed in [27]: accuracy(Ac), sensitivity(Se), specificity(Sp) and precision(Pr). All four metrics are calculated based on true positive TP, false positive FP, false negative FN and true negative TN in a binary confusion matrix. Therefore all four metrics are calculated for each class by converting the 4x4 matrix to a 2x2 matrix.

While cumulated classification results are demonstrated in Table 4.1, the robustness of the proposed method should be evaluated based on the performance variation over 22 test records in DS2. Hence medians and IQRs(interquartile range) for each metric and each class are included in Table 4.2 to represent the robustness of proposed methods. The lower variation between performances measured on different tapes, the more robust the system is. In Table 4.2, we observe that among all abnormal classes, the proposed algorithms demonstrated stable performance on class V and high performance but less stable on class V and F.

As MITDB is widely used to verify ECG classifier performance, we compared the proposed system with five significant methods proposed in literature. According to AAMI standards, ECG classifier performance should be evaluated over the binary classification performance

Table 4.3: V and S classification performance compared with five algorithms in literature using 11 common records in MITDB

Methods		VEB		SVEB			
Methods	Ac	Se	Sp	Ac	Se	Sp	
Proposed	96.6	98.2	92.4	98.63	88.89	99.41	
Hu et al. [27]	94.8	78.9	96.8	N/A	N/A	N/A	
de Chazal et al. [33]	96.4	77.5	N/A	N/A	N/A	N/A	
Jiang and Kong [29]	98.8	78.9	96.8	97.5	74.9	98.8	
Ince <i>et al.</i> [30]	97.9	90.3	98.8	96.1	81.8	98.5	
Kiranyaz et al. [31]	98.9	95.9	99.4	96.4	68.8	99.5	

of Ventricular (VEB) versus non-VEB types and Supraventricular (SVEB) versus non-SVEB types. For methods proposed in literature, same evaluation metrics are deployed on records from MITDB. To standardize the metrics, we select 11 common ECG records from all 5 methods and compared the median of each classification metrics over these 11 records. The comparison results are demonstrated in Table.4.3. Generally speaking, the proposed method shows higher sensitivity for both VEB and SVEB. Especially for SVEB, the proposed method has advantage on all three metrics over other five methods in the literature.

4.6.2 Prediction Performance

As an important characteristic of the proposed methods, yellow alarms triggered by Personal Classifier after feature space reshaping indicate a higher probability of observing subsequent abnormalities. In order to verify this functionality, all beats following a yellow alarm is investigated for each yellow alarm. Abnormality type which occurs the earliest within the window is recorded. Similar to confusion matrix for classification evaluation, the performance of prediction can be summarized by a confusion matrix with the 3 abnormal types. Probabilities of observing an certain type of abnormal beats after a yellow alarm is calculated using the prediction confusion matrix and compared to the prior probability of observing the same

Table 4.4: predictive probability versus prior probability without windowing

		# of	predi	cted ground truth	% of predicted ground truth			
	V	S	F	V	S	F		
	V	467	122	14	77.45	20.23	2.32	
yellow alarm	S	36	15	0	70.59	28.41	0	
	F	40	60	5	38.10	57.14	4.76	
total		543	197	19	71.54	25.96	2.50	

type of abnormality. This process is formulated in the following two equations:

$$P(\hat{y}_{k+i} = X_r | \hat{y}_k = X_y) = \frac{\text{\# of } y_{k+i} = X \text{ after } \hat{y}_k = X_y}{\text{\# of true alarms after } \hat{y}_k = X_y}$$

$$P(\hat{y}_{k+i} = X_r) = \frac{\text{\# of true alarm of type } X (y_k = X)}{\text{\# of all true alarms}}$$
(4.10)

The capacity of predicting each type of abnormalities is evaluated by comparing $P(\hat{y}_{k+i} = X_r | \hat{y}_k = X_y)$ and $P(\hat{y}_{k+i} = X_r)$. A shown in Table.4.4, the probability of observing a certain type of abnormalities after a yellow alarm are higher than its prior for each type of abnormalities. For example, without knowing the type of a yellow alarm, the probability of observing a type V sample is 71.54% while the probability of observing a type V sample given a type V yellow alarm was triggered is 77.45%. The improvement are consistent among all three types of abnormalities but the system has stronger predicting capacity for type S. In order to study the time window in which real abnormality occurs after yellow alarms, we also studied a window of 10 consecutive samples following a yellow alarm. Similarly, prior and posterior probabilities are compared to evaluate the performance as shown in Table.4.5.

Compared with the result without windowing, the predicting performance within 10 beats window shows that the proposed algorithm can better predict the occurrence of abnormalities in a centrain time window. Especially for type S, the probability of observing a type S sample

Table 4.5: predictive probability versus prior probability within 10 beats' window

	# of	predi	cted ground truth	% of predicted ground truth			
	V	S	F	V	S	F	
	V	290	85	12	74.94	21.96	3.10
yellow alarm	S	22	13	0	62.86	37.14	0
	F	29	37	6	40.28	51.39	8.33
total		341	135	18	69.03	27.32	3.64

within 10 beats after a yellow alarm is 27.32% while given that the yellow alarm is type S, the posterior probability is raise to 37.14%. With almost 10% increase, it's proved that the yellow alarm types are informative. The results shows that same improvements are made within the 10-sample window as well. In general, the predicting performance are promising, indicating the efficiency of personal classifier and deviation analysis.

Chapter 5

Conclusions And Future works

5.1 Conclusions

In this thesis, we a patient-adaptable ECG classification framework. The system has a two-staged hierarchical classifier structure including Global Classifier and Personal Classifier. While Global Classifier is designed to filter the signal with severe distortion and abnormal waveforms by triggering red alarms and pass other samples to the deviation detection stage. In this stage, the personal dynamic normal cluster is constructed and used to specify the normal range for each patient. By comparing the current sample and personalized normal range, this module decides if a yellow alarm will be triggered to provide predictive information about upcoming abnormalities. If a sample is detected with deviation towards abnormal clusters, it will be passed to the Personal Classifier and labeled as one of the three abnormal types. Whereas samples without deviation are further feed back to personal dynamic normal cluster to update the classification system about the newest personal normal range.

In Chapter 3, a kernel based nonlinear transformation is proposed to address the problem

of cluster topology in original feature space. Inspired by Support Vector Machine, kernel functions are deployed in this method as a spatial transformation function. The target topology is formulated as two objective functions so that by tuning the parameters in kernel function, the system is able to select the best transformation for the following predicting stage. This non-convex multi-objective optimization is solved with Multi-Objective Particle Swarm Optimization. In order to validate improvement by using high order kernel function, we compared the Pareto front generated with linear combination of original features and mapped high order features with polynomial kernel. The result verifies that applying high order kernel function allows more degree of freedom so that the topology can be further optimized according to objective functions. Having this concept proved, we applied this method on MITDB test data and obtained similar sensitivity and specificity as proposed in the literatures. More importantly, the predicting capacity of yellow alarms are analyzed. The performance is quantified by comparing prior probability and posterior probability giving the types of yellows alarm. The comparison result shows that a promising improvement has been made by applying the nonlinear transformation.

While the method in Chapter 3 demonstrated capacity of predicting upcoming abnormalities, it's challenging to interpret the mechanisms of the systems and thus hindering the generalization of predictive warning to other applications of biomedical signals. Therefore, the main object of Chapter 4 is developing a classification system with abnormality predicting capacity based on spatial topology studied in Chapter 3. In Chapter 4, we proposed a novel spatial transformation specifically designed to reshape the feature space according to angles between cluster center. In this method, between cluster cosine distance are optimized through orthogonalization in spherical coordinate space and within cluster variance is reduced by a mapping function which is fitted piecewise with a basis function. The basis function proposed in the chapter has the feature of saturating at the boundaries, similar

to sigmoid function but more flexible. An advantage of deploying such basis function is that the cluster geometry may be preserved after spatial transformation. With this novel module integrated in the patient-adaptable classification framework, the performance of this system is evaluated through classification and prediction results on the test set data. The classification results show that by triggering yellow alarm through this method, specificity of abnormal types is improved. Especially for Supraventricular types, the proposed system performs better than all 5 methods in the literature. The same conclusion holds for prediction performance. Compared to the method proposed in Chapter 3, this method improves predicting capacity for all abnormal classes and the most significant improvement is for type S. Moreover, we also studied the time delay of real abnormalities following a yellow alarm. It's proved that most of the real abnormalities occurs within 10 beat after a yellow alarm. Generally speaking, the system is proved to be efficient both in classification and prediction.

5.2 Future works

In this research, we focused on two challenges of ECG classification, namely, inter-patient variation and anomaly prediction. The framework of patient-adaptable classier includes both features. The methods of improving prediction accuracy are proposed and studied. While the result shows the efficiency of designed system, further improvements can be made through research. The following tasks can be resolved as a continuation of this research:

- Research on other kernel functions which is potentially a better transformation for spatial topology optimization.
- Investigate on the deterministic solution for the objective functions proposed in Chapter 3.

- Assess the performance of proposed spatial transformation on other biomedical signals with similar characters as ECG signal
- Improve the deterministic mapping function in Chapter 4 by including the variance of individual clusters into function parameters

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