# HEART MURMUR DETECTION AND ANALYSIS USING MULTIPOINT AUSCULTATION SYSTEM

KAMARULAFIZAM BIN ISMAIL

UNIVERSITI TEKNOLOGI MALAYSIA

# HEART MURMUR DETECTION AND ANALYSIS USING MULTIPOINT AUSCULTATION SYSTEM

### KAMARULAFIZAM BIN ISMAIL

A thesis submitted in fulfilment of the requirements for the award of the degree of Doctor of Philosophy (Biomedical Engineering)

Faculty of Biosciences and Medical Engineering Universiti Teknologi Malaysia

AUGUST 2015

Dedicated to my beloved wife Azlin Abd Jamil, my beloved children, Dania Sofea, Danny Iskandar, Daniel Akashah and Diana Maisara, and my beloved father, mother, brothers, sisters & friends.

#### ACKNOWLEDGEMENT

First of all, I wish to thank heartedly my supervisor Professor Ir. Dr. Sheikh Hussain Bin Shaikh Salleh. He is the one who introduced me to the field of biomedical signal processing research. It is his endless guidance of technical knowledge and methods of conducting research that built my foundation on heart sound analysis field. His moral and financial support survives me throughout my PhD study. His attitude and enthusiasm in conducting research and his consistent vision to make the research an important asset for the country and the next generation always inspire me.

Secondly, I am thankful to the Director for Centre for Biomedical Engineering, Prof. Dato' Ir Dr Alias Mohd Nor for his continuous encouragement and support. And to all of my research colleagues at the Center of Biomedical Engineering (CBE), namely Dr Ting Chee Ming, Ahmad Kamarul Ariff, Arief Ruhullah, and others. Thank you very much for your generosity to share the resource and knowledge with me. Special thanks to CBE for providing the resource and conducive environment to conduct my research.

Finally, I wish to thank my family and friends for their support. I am especially grateful for my parents for all their sacrifices in upbringing me. The encouragement and support for me in pursuing my research career are appreciated sincerely. I would like to thank my wife Azlin, she always by my side, comfort me when I am discouraged, take care of me when I am busy, and share my happiness when I delighted.

### ABSTRACT

The study of phonocardiogram (PCG) in diagnosing valvular heart disease has gathered increasing attention over the past few years. Heart sound auscultation is performed at the primary care center by physician and the results are subjected to the skills and hearing ability. This has caused unnecessary referral and send home subject with potential heart disease. This issue has led to the establishment of standardized and computerized system to analyze the heart sound. This thesis investigates the optimal approach in establishing a reliable system to acquire and process heart sound to differentiate between normal and abnormal pattern. Previous studies are based on the analysis using heart sound that is recorded from single stethoscope which provides limited information regarding the heart disease. In this study, the recording based on four stethoscopes is used to record sound from four different valves with optimized analog instrumentation design. Beamforming algorithm is utilized to localize the actual source of the disease sound from all of the four recorded sound by focusing with respect to the angle of arrival of the desired disease signature. It is then followed by the implementation of Time Frequency (TF) algorithm with optimal Extended Modified B-Distribution (EMBD) kernel to suppress noises, analyze and represent the features. The experiments were conducted utilizing PCG signal that was recorded from real subject from Hospital Sultanah Aminah Johor Bahru. Each subject was screened by an echocardiogram machine. The disease was confirmed by cardiologist before the PCG recording procedure was performed. The result shows significant improvement in the quality of information that is preserved in the beamformed signal. The suggested framework is able to improve the heart murmur detection rate up to 95%. In conclusion, the localization of the exact location of the diseased sound has helped to improve the disease detection accuracy based on multi-point heart sound diagnostic system.

### ABSTRAK

Kajian tentang bunyi degupan jantung untuk menganalisa penyakit jantung telah mendapat perhatian yang semakin meningkat sejak beberapa tahun yang lalu. Pengambilan bunyi jantung dilakukan di pusat penjagaan kesihatan oleh ahli fisiologi dan keputusan analisa adalah subjektif mengikut tahap kemahiran dan pendengaran ahli fisiologi tersebut. Keadaan ini menyebabkan pesakit yang tidak mempunyai penyakit dirujuk kepada pakar dan pesakit yang berpotensi berpenyakit dihantar pulang. Isu ini telah membawa kepada pengwujudan sistem berkomputer yang selaras untuk menganalisa isyarat bunyi jantung. Tesis ini mencari kaedah yang optima bagi mewujudkan sistem yang berkebolehpercayaan yang tinggi untuk memproses isyarat bunyi jantung untuk membezakan bunyi jantung normal dan tidak normal. Kajian sebelum ini menunjukkan pelbagai eksperimen telah dilakukan tetapi hanya berdasarkan bunyi yang dirakam dari satu lokasi dan mempunyai maklumat yang terhad. Kajian ini mencadangkan rakaman isyarat menggunakan sehingga empat stetoskop dengan rekabentuk instrumentasi yang optima. Kaedah pembentukrasuk digunakan untuk mengenalpasti sumber bunyi asal dengan memfokuskan kepada arah bunyi jantung. Ianya kemudian diikuti oleh penggunaan teknik frekuensi-masa menggunakan algoritma modifikasi B-Distribution lanjutan untuk mengurangkan hingar, menganalisa isyarat dan mewakilkan ciri penting isyarat. Eksperimen dijalankan menggunakan data dari Hospital Sultanah Aminah Johor Bahru. Setiap subjek diperiksa terlebih dahulu menggunakan mesin *echocardiograph*. Jenis penyakit disahkan dengan bantuan pakar jantung. Hasil kajian mendapati kualiti isyarat bunyi menjadi lebih baik selepas di proses oleh teknik pembentukrasuk. Keseluruhan rangka kerja ini mempu meningkatkan keupayaan mengecam bunyi desiran jantung sehingga 95%. Kesimpulannya, kebolehan menentukan sumber bunyi yang tepat telah membantu meningkatkan ketepatan pengesanan bunyi jantung berdasarkan sistem diagnostik isyarat jantung pelbagai titik.

# TABLE OF CONTENTS

CHAPTER

1

## TITLE

PAGE

DECI	LARATION	ii	
DEDI	ICATION	iii	
ACK	ACKNOWLEDGEMENT		
ABST	TRACT	v	
ABST	TRAK	vi	
TABI	LE OF CONTENTS	vii	
LIST	LIST OF TABLES		
LIST OF FIGURES			
LIST OF ABBREVIATIONS			
LIST OF SYMBOLS			
LIST OF APPENDICES			
INTRODUCTION			
1.1	Introduction	1	
1.2	The Organization of the Thesis	2	
1.3	Problem Background	3	

1.4	Problem Statement	5
1.5	Objectives	5
1.6	Scope	6
1.7	The Contribution of the Study	7

## LITERATURE REVIEW

9

2.1	Introduction		
2.2	Cardiac Auscultation Proficiency Trends		
2.3	Principles of Cardiac Auscultation		
2.4	Auscultory Sites		
2.5	The Cardiac Cycle- Sound and Murmurs	14	
2.6	Clinically Important Cardiac events	16	
	2.6.1 Early Systolic Ejection Click	16	
	2.6.2 Mid Systolic Click	16	
	2.6.3 Opening Snap	17	
	2.6.4 Third Heart Sound (S3)	17	
	2.6.5 Fourth Heart Sound (S4)	17	
	2.6.6 Pericardial Rub	18	
	2.6.7 Aortic Stenosis (AS)	18	
	2.6.8 Aortic Insufficiency (AI)	19	
	2.6.9 Mitral Stenosis (MS)	20	
	2.6.10 Mitral Regurgitation (MR)	21	
	2.6.11 Patent Ductus Arteriosus (PDA)	21	
2.7	The Recording using Electronic		
	Stethoscope	22	
2.8	Sound Localization	24	
2.9	The Acquisition Apparatus	29	
2.10	Denoising	31	
2.11	The Segmentation Issues	32	
2.12	Biosignal Analysis of Heart Murmurs	35	
2.13	Automatic Classification of Heart Sound	41	
2.14	Alternate Method for Heart Sound Analysis	43	
2.15	Beamforming Method	44	
	2.15.1 Delay and Sum Method	46	
	2.15.2 Microphone Array Design	47	
2.16	Summary	50	

# THE DESIGN ACQUISITION APPARATUS 52

3

4

3.1	Introduction	
3.2	Single Location Recording	53
3.3	Design Requirement and Philosophy	54
3.4	The Design of Heart Diagnostic System	55
	3.4.1 The Transducer	56
	3.4.2 The Analog Front-End	60
	3.4.3 Computer Interfacing	63
3.5	Single Supply 5V Data Acquisition	
	System	65
3.6	Dual Supply 12V Data Acquisition	
	System	74
3.7	Summary	76

## THE PROCESSING FRAMEOWRK 77

4.1	Introduction		77
4.2	Database		78
4.3	Patient Preparation		79
4.4	Stethoscope Positioning and Placement		80
4.5	Signal Examination and Segmentation		80
4.6	The Sound of Interest.		83
	4.6.1	The Clinical Significance of S1	
		and S2	84
	4.6.2	The Clinical Significance of	
		Murmurs	89
4.7	Sound	Localization using Beamforming	89
	4.7.1	Beam Pattern for Source	
		Localization	92
	4.7.2	Spatial Filtering	95
4.8	The T	ime Frequency Analysis	98
4.9	Summ	nary	104

## **RESULTS AND DISCUSSION**

5.1	Introd	uction	105
5.2	The Design of Acquisition System		
5.3	Instru	mentation	106
5.4	Morpl	nological Analysis	111
5.5	Multi-	location Heart Sound Recording	115
5.6	Comp	arison between the Proposed Designs	
	and W	elch Allyn System	117
5.7	Analy	sis on S1 and S2	118
5.8	Heart Sound Localization using		
	Beam	forming	121
	5.8.1	Case 1	121
		5.8.1.1 Time Delay Calculation	
		Method	122
		5.8.1.2 Delay and Sum	
		Beamforming Method.	126
	5.8.2	Case 2	129
		5.8.2.1 Time Delay Calculation	
		Method.	129
		5.8.2.2 Delay and Sum	
		Beamforming Method.	131
	5.8.3	Case 3	134
		5.8.3.1 Time Delay Calculation	
		Method	134
5.9	Time-	Frequency Analysis	135
5.10	Summ	nary	145

# 6 CONCLUSIONS AND FUTURE WORK 146

6.1	Conclusion	146
6.2	Future Work	148

REFERENCES
------------

Appendices A-D

165-180

150

# LIST OF TABLES

<b>TABLES</b>	NO.
---------------	-----

## TITLE

## PAGE

3.1	Sensor design matrix	60
3.2	Specification of transducer	60
3.3	Comparison of commonly used operational amplifier	67
4.1	Types of beamforming	90
5.1	Digital value representing uptrend for 12V and 5V	
	System	114
5.2	Digital value representing sideways movement for 12V	
	and 5V system	115
5.3	Performance comparison between Welch Allyn	
	Stethoscope system and 100 Analyzer system.	118
5.4	The peak difference between the heart sound signal's	123
	peak and the ECG signal's peak from subject 1.	
5.5	Time delay between all four stethoscopes	123
5.6	The difference in distance	124
5.7	The peak difference between the heart sound signal's	129
	peak and the ECG signal's peak from subject 2.	
5.8	Time delay between all four stethoscopes	130
5.9	The difference in distance	130

# LIST OF FIGURES

FIGURES NO	D. TITLE PA	PAGE	
2.1	Auscultory areas	14	
2.2	Complete cardiac cycle	15	
2.3	Cardiac events.	15	
2.4	Components typically occurs mid diastolic, late diastolic and mid	l	
	systolic	18	
2.5	Crescendo decrescendo murmur	19	
2.6	Decrescendo murmur	19	
2.7	Crescendo murmur	20	
2.8	Mitral stenosis murmur	20	
2.9	Mitral regurgitation murmur	21	
2.10	Continuous murmur	22	
2.11	Typical system setup for automatic heart sound diagnostic system	n 23	
2.12	QRS complex with R point to determine every cycle of the one		
	minute data of the heart sound	23	
2.13	Echocardiogram image	25	
2.14	Planar waveform reaching linear microphone array	46	
2.15	Beam shape	48	
2.16	Cardiac image constructed different beamforming technique	49	
2.17	Delay and sum beamformer	50	
3.1	Auscultation Location	53	
3.2	Overall system design of electronic processing interface	55	
3.3	Sample waveform of heart sound	59	
3.4	Operational amplifier using INA118	61	
3.5	The cascaded filter and amplifier for heart sound acquisition		

	System	63
3.6	The USB interface circuit design for heart sound acquisition	
	System	65
3.7	CMRR for INA118	68
3.8	CMRR for LF412	69
3.9	CMRR for OPA330	69
3.10	CMRR for OP07	69
3.11	CMRR for LF347	70
3.12	CMRR for AD620	70
3.13	Rescaled design of heart sound and ECG data acquisition system	
	using 3.3V supply system	71
3.14	Heart sound recording using Welch Allyn and the proposed	
	device for a normal subject	73
3.15	Information lost due to improper gain setting and small range	
	operating voltage	74
4.1	Mobile medical trolley to carry recording device	79
4.2	Heart sound signals	82
4.3	Heart sound and ECG timing characteristics	83
4.4	Effect of inspiratio and expiration upon the morphology of S1	
	and S2	84
4.5	Location of each electronic stethoscope	90
4.6	Beamforming flowchart	90
4.7	Overall beamforming processing pathway	91
4.8	Source localization simulation using 400Hz source and four	
	electronic stethoscope array	93
4.9	Comparison of beam width versus the number of electronic	
	stethoscope	94
4.10	Comparison of beam width versus frequency using two electronic	
	stethoscope	94
4.11	Decibel scale plot of a four electronic stethoscope array	
	beam pattern for a 600Hz signal at 90 degree	95
4.12	The noise sources at 300Hz attenuated at about 3dB using two	
	electronic stethsoscopes separated at 60 degrees	96
4.13	An array of two electronic stethoscope with sources	

	seperated by 90 degrees with attenuate noise by 13dB.	96
4.14	An array of four electronic stethoscope with sources	
	seperated by 60 degrees with attenuate noise by 12dB	97
4.15	An array of four electronic stethoscope with sources	
	seperated by 90 degrees with attenuate noise about 12dB	
	lower than the source signal	97
5.1	The constructed recording apparatus	106
5.2	Signal recording for 5V system and 12V system	107
5.3	Three cycles of close-up signal showing details of raw	
	ECG signals	108
5.4	One cycle zoomed version. (a) Distorted signal using	
	5V system, (b) 12V system	109
5.5	QRS complexes. (a) Distorted QRS complex of 5V system, (b)	
	Smooth peak of QRS complex for 12V system	110
5.6	Position of actual peak and distorted peak, (a) Peak of	
	5V system, (b) Peak of 12V system.	110
5.7	Signal peak delay difference between 5Vsystem and	
	12V system	111
5.8	Effect of different voltage range on signal morphology	112
5.9	Effect of different voltage range on signal morphology	112
5.10	Tracking of signal changes	113
5.11	Design of stethoscope housing	116
5.12	Transducers arrangement as a recording system	117
5.13	Amplitude comparison for S1 and S2 for location Aortic (V1),	
	pulmonic (V2), tricuspid (V3) and mitral (V4)	119
5.14	Amplitude comparison for S1 and S2 for location Aortic (V1),	
	pulmonic (V2), tricuspid (V3) and mitral (V4)	119
5.15	Amplitude comparison for S1 and S2 for location Aortic (V1),	
	pulmonic (V2), tricuspid (V3) and mitral (V4)	120
5.16	Amplitude comparison for S1 and S2 for location Aortic (V1),	
	pulmonic (V2), tricuspid (V3) and mitral (V4)	120
5.17	Coordinates system for four location stethoscopes	122
5.18	Coordinates of unknown sound source form data 1	125
5.19	Calculated unknown sound of the source at (-38, 61).	125

5.20	Heart sound signal data 1 from subject 1 at four location	126
5.21	Original signal and received signal with beamforming method	127
5.22	Angle of arrival calculation method	128
5.23	Arrival angle (32 degree)	128
5.24	Coordinates of unknown sound source form data 2	130
5.25	Calculated unknown sound of the source (-43, 58)	131
5.26	Heart sound signal data 2 from subject 2 at four locations	132
5.27	Original signal and received signal with Beamforming	132
5.28	Angle of arrival calculation method	133
5.29	Arrival angle	133
5.30	Heart sound signal recorded from normal subject at four	
	locations.	134
5.31	Source of sound for the normal heart sound	134
5.32	EMBD plot for normal heart sound	136
5.33	EMBD plot for an abnormal heart sound	137
5.34	Multi-location Time-Frequency representative of normal	
	heart sound before and after Beamforming .	141
5.35	Multi-location Time-Frequency representative of abnormal heart	
	sound before and after Beamforming.	144

# LIST OF ABBREVIATIONS

A2	-	Aortic Valve Closure Sound
ADC	-	Analog to Digital Converter
AI	-	Aortic Insufficiency
ANN	-	Artificial Neural Network
CHF	-	Coronary Heart Failure
CI	-	Cochlear Implant
CMRR	-	Common Mode Rejection Ratio
CWT	-	Continuous Wavelet Transform
DFT	-	Discrete Fourier Transform
DSBF	-	Delay and Sum Beamforming
DTW	-	Dynamic Time Warping
E	-	Energy
ECG	-	Electrocardiogram
EMD	-	Empirical Mode Decomposition
EMBD	-	Extended Modified B-Distribution
FFT	-	Fast Fourier Transform
GP	-	General Practitioner
HA	-	Hearing Aid
HMM	-	Hidden Markov Models
ICS	-	Intercostal Space
IDFT	-	Inverse Discrete Fourier Transform
LPC	-	Linear Predictive Coding
LPCC	-	Linear Predictive Coding Cepstrum
LSB	-	Left Sternal Border
M1	-	Mitral Valve Closure Sound
MR	-	Mitral Regurgitation

MS	-	Mitral Stenosis
MBD	-	Modified B-Distribution
MCE	-	Minimum classification error
MFCC	-	Mel-Frequency Cepstral Coefficients
MFPC	-	Mel-Frequency Power Cepstrum
MLP	-	Multi-Layer Perceptron
MVP	-	Mitral Valve Prolapse
NN	-	Neural Network
OS	-	Opening Snap
P2	-	Pulmonic valve closure sound
PCG	-	Phonocardiogram
PLP	-	Perceptual Linear Prediction
PR	-	Pulmonary Regurgitation
PS	-	Pulmonary Stenosis
PVWD	-	Pseudo Wigner Ville Distribution
RSB	-	Right Sternal Border
S1	-	First Heart Sound
S2	-	Second Heart Sound
<b>S</b> 3	-	Third Heart Sound
S4	-	Fourth Heart Sound
SA	-	Sinus Atria
SNR	-	Signal to Noise Ratio
STFT	-	Short Time Fourier Transform
SVM	-	Support Vector Machine
T1	-	Tricuspid Valve Closure Sound
TF	-	Time-Frequency
TFD	-	Time-Frequency Distribution
TR	-	Tricuspid Regurgitation
USB	-	Universal Serial Bus
VSD	-	Ventricular Septal Defect
WT	-	Wavelet Transform

# LIST OF SYMBOLS

$\Delta_{\mathrm{m}}$	-	Delay of the m <sup>th</sup> sthethoscope
$\gamma_m(i)$	-	Scalar representation of stethoscope amplitude
τ	-	Signal delay
$S(\omega)$	-	Source signal from stethoscope
$Z(\omega, \theta)$	-	Signal mixture of all stethoscopes
$H(\omega, \theta)$	-	Transfer function for source signal
z(t)	-	Time domain representative of signal
E(t)	-	Energy representative of signal
S(f)	-	Frequency domain representation of signal
$\rho(t,f)$	-	Time frequency representation of signal
$G(t, \tau)$	-	Time lag kernel
g(ν, τ)	-	Doppler lag kernel function
β	-	Smoothing variable for time-frequency

# LIST OF APPENDICES

APPENDIX	TITLE	
А	Ethics committee approval from the Ministry of Health	165
В	Patient Consent Form	169
С	Patient Information Sheet	172
D	Publications	179

### **CHAPTER 1**

### **INTRODUCTION**

### 1.1 Introduction.

Heart disease is the number one killer disease in most countries in the world (Nichols M., 2014). The statistic shows significant increment in mortality rate each year. Regardless of the causes, most fatality is caused by the inability to detect this disease at the early stage (Michael S., 2006). Detection at the earlier stage could save many lives and able to reduce the treatment cost tremendously.

Human heart exhibits a plethora of information regarding its health status and working condition via its electrical signal known as electrocardiogram (ECG). The acoustic signal generated is known as phonocardiogram (PCG) or heart sound. The ECG is an electrical impulse originating from Sinoatrial (SA) node as an effect of polarization and depolarization of heart tissue (Pipberger *et al.*, 1962). This electrical impulse initiates the working mechanism of blood pumping activity, which open and close four valves in the heart and produces the PCG.

ECG has been used for more than a century in diagnosing heart disease (Pipberger *et al.*, 1962). The information contain in ECG signal is more related to the

heart tissue conduction issues and the pumping regulations of the valves. However it does not describe the pumping capability of each valve, the condition of each valve and the overall efficiency of the heart. Since heart sound is capable in providing such information, it has been used as a primary screening tool together with ECG when diagnosing patients with suspected heart disease. In general practice, a physician will perform heart sound auscultation before recommending ECG screening. This procedure makes complete sense to examine the valve first then ECG on the basis that ECG is the one which regulates the valves operation. Any discrepancy on the valve operation may possibly be caused by the ECG.

### **1.2** The Organization of the Thesis.

This thesis is divided into 6 chapters. Chapter 1 introduces the issues and related motivational element, which drives the need to perform this research. It also covers the research objectives as well as outlining the research limitation, bridging the research gap and scope. Chapter 2 describes the literature review on anatomy of heart sound and it's relation to heart disease. The significant heart sound signatures of each disease are explained which is important in extracting unique features. It is then followed by a review of heart sound analysis. A number of heart disease detection methods are elaborated in terms of its strength and weakness and how the proposed method emerged. Chapter 3 describes the design and development of 5 channels acquisition apparatus that is used in this research. Chapter 4 describes the processing procedure using beamforming and time-frequency analysis. In chapter 5, the results are presented with related discussions. In chapter 6, the conclusions are presented.

### **1.3** Problem Background

Developing the skill of listening to the heart sound or also known as auscultation requires years of training. This ability is different from one physician to another. The outcome of interpretation is also subjective. A physician is required to be trained regularly in order to maintain the auscultation skill (Tavel, 1996 and Cheitlin *et al.*, 1997). The traditional acoustic bell shape stethoscope is capable in delivering sound from 100Hz to 200Hz. However, most of the heart sounds frequency content lies at the lower frequency band, which is as low as 50Hz to 500Hz (Abbas, 2009). This provides limited heart sound information and led to many false diagnoses resulting in numerous unnecessary referrals. The subject with heart disease is sent back when the disease is still undetected. Many studies have proven that as many as 87% of patients that are referred to cardiologists are as a result of false alarms (Pease, 2001).

Since the introduction of echocardiography technology which is based on ultrasound imaging, it has become a gold standard in heart disease verification. Here the heart sound and ECG are only used as pre-screening tools. However, the implementation is restricted by the availability of this tool due to high acquisition cost. A typical machine would cost up to 1.5 million ringgit. Only big hospitals can afford the cost and only a few units can be made available. This limited number of machine could not be a solution to help the large number of the population. As a result, the patients have to wait for the disease to be diagnosed before it is confirmed and treated. Such machine has been around for more than a decades and heart disease still remain as the number one killer disease. Therefore, echocardiogram seems not to be the solution to current scenario. A much cheaper machine with the capability to detect heart disease from the very earlier stage is critically needed. This is the main subject of this thesis in order to address the issue raised earlier in heart related disease. Diagnosing heart disease based on heart sound will require an ECG signal to be recorded together and displayed side by side. This will help the physician to determine the beginning of a cardiac cycle. However, it is rather difficult to find a system that records heart sound and ECG simultaneously. A typical system that is available in the medical field is either to record the heart sound or just ECG. Even if there is, the principle behind the design is just for the sake of monitoring and not tailored for heart disease diagnosis. Diagnosing bio-signal requires high precision data with specialized design of analog and digital circuitry that preserve not only the information but also remove the unwanted noise which is one of the concerns of this study.

Since the heart is operated by four valves, and typically diagnosed down to each valve, it makes perfect sense to listen to all of the sound that is produced by each of these valves. Manual diagnosis performed by physician usually moves the stethoscope around the chest area to find abnormal sound produced by the valves. Once the location is identified, the physician listens closely and starts to list down several suspected diagnosis based on the sound. Automated diagnosis would require all four locations of heart sound in order to be able to locate the actual source of the problem. Recording one after another will not help to locate the problem in real-time, thus simultaneous recording is suggested in this thesis.

### **1.4 Problem Statement**

The study is motivated by the need of solutions from the following problems:

- Lack of fast and reliable screening tool to aid the general practitioner (GP) in the primary care center (echocardiogram machine cannot be place in all the clinics).
- The use of single stethoscope to acquire heart sound provides limited information. A tool that is able to maximize the information acquisition from the beating heart, down to each and individual valve is critically needed. The multi-point auscultation device which records four sounds from four valves simultaneously with lead II ECG provides massive advantages.
- The correct information has to be extracted from the heart sound from the right location on the chest. Recording four heart sound simultaneously provides localization advantage. An efficient algorithm to pin point the exact location of murmur is necessary to improve detection.
- The general practitioner needs support in making decision. A reliable and accurate scientific presentation and visualization would be of a great advantage in deciding whether a subject should be referred or otherwise.

#### 1.5 Objectives.

In this thesis, the research objective is concerned with identifying murmurs based on heart auscultation. In particular, the thesis focuses in the improvement of system hardware design by specific development of the multi-point bio-signal input. Various performance measures are used to evaluate the beamforming auscultation system for different aspect of performance. The results presented reflect the acceptable level of initial performance of the system. The research objectives of this study are as follows:

- To design and develop a 5 channel data acquisition system for the heart sound and ECG.
- To perform multi-point auscultation to acquire four heart sound simultaneously.
- To enhance the beamforming auscultation system for heart murmur analysis.
- To evaluate the performance of time-frequency analysis of heart murmurs.

#### 1.6 Scope.

The main concern of this study is to design a new five channel data acquisition system for multi-point auscultation of the heart sound. The primary focus is on the design of the new hardware for signal acquisition as the available data processing system are only capable of monitoring of the heart sound. A special emphasis is placed on the evaluation methods with real microphone recording involving simultaneous heart sound signals, as opposed to computer-generated simulation. This new design will be followed by an introduction to a new procedure to process the multi-channel heart sound which enables the localization of heart murmurs utilizing beamforming algorithm. As the beamforming is usually used in communication, this is the first time to utilize the approach in biomedical signal particularly in heart sound.

There are several approaches to time frequency analysis (Cohen *et al.*, 2001) which can be used to tackle this problem, but this does not come to focus as the extended modified B-distribution algorithm is used here. The modification is necessary to fit the nature of heart sound signal model, which is generated by vibration collected by microphone.

The scope of the study is limited on these specific issues:

- Ensure the proposed 5 channels design is capable of acquiring high quality bio-signal data, which correlates the ECG and heart sound signal.
- Utilization the beamforming algorithm to localize heart murmurs based on multi-point auscultation system.
- Utilize the extended modified B-distribution algorithm to visualize the presence of murmurs.

#### **1.7** The Contribution of the Study.

In this research, an optimal method of accessing cardiac abnormalities is deployed. There are several major contributions that have been achieved from this research as follows:

- A new 5-channel analog front-end system is developed to ensure optimal signal quality is acquired. The consideration started from the selection of proper transducer which is sufficiently sensitive to capture the vibration of beating heart from the human chest. The instrumentation stage is carefully designed to ensure all possible information is preserved with the most minimum information losses. The selection of operational amplifier, filter, analog to digital converter, the operating voltage is discussed in detail in chapter 3.
- This study proposes multi-point auscultation technique in acquiring the heart sound. Typically, heart sound is acquired at one location and disease is determined using that information. As the acquired sound originated from four locations namely aortic valve, pulmonic valve, tricuspid valve and mitral valve, and the disease is also associated to each and individual valve, it makes perfect sense to acquire all the four sound at the same time and use the combination of all the sound as input to the processing stage. This could provide more information about the dynamic operation of all valves especially when it comes to diseases.
- A physician usually start listening to heart sound at a position and move the stethoscope around until the desired diseased sound is audible. This justify that disease sound is not always present at the location where it is originally produced which are the valves. The sound has to be mapped out around the valves. To adapt this approach, beamforming technique is used to identify the actual source of the sound. It is hypothesized that beamforming method is able to highlight the important sound from all the given four valves sound and to pin point the location that heart sound should be acquired.
- Time-frequency analysis is a popular tool to visualize signal content in term of energy, time and frequency. It is usually derived from communication research and application. Time frequency analysis utilizing B-distribution algorithm is modified to fit medical application. This could provide an improved presentation of heart sound and murmur.

#### REFERENCES

- Abbas AK, Bassam R (2009), *Phonocardiology Signal Processing*, Morgan & Claypol Publisher.
- Alim O.A., Hamdy N., and El-Hanjouri M.A. (2002), Heart diseases diagnosis using HMM, in *Proc. National Radio Science Conference (NRSC)*, pp. 634-640.
- Amit G., Noam Gaviriety, Nathan Intrator (2009), Cluster analysis and classification of heart signal, *Biomedical Signal Processing and Control 4*, pp. 26-35.
- Ashish H, Sushil T, Panse M. S (2011), Digital Stethoscope for Heart Sounds, 2nd International Conference and workshop on Emerging Trends in Technology (ICWET).
- Azra'ai, R.A., bin Taib, M.N., Tahir, N.M. (2008): 'Artificial neural network for identification of heart problem'. Proc. Second Int. Conf. on Signal Processing and Communication Systems, Gold Coast, Australia, pp. 1 – 6
- Avants B. and Gee J. (2003), Continuous curve matching with scale-space curvature and extrema-based scale selection, in *Proc. Scale-space Methods in Computer Vision*, p.1079, 2003.
- Bahadirlar Y. and Gulcur H. O. (1998), Cardiac passive acoustic localization: cardiopal, *Turkish Journal of Electrical Engineering and Computer Science*, Turkey, vol. 6, pp.243–259.
- Bahadirlar Y. and H. Gulcur O. (2001), Time-frequency cardiac passive acoustic localization, 23th IEEE International Conference on Engineering in Medicine and Biology, Turkey, vol. 2, pp.1850–1853.
- Balster DA, Chan DP, Rowland DG, Allen HD (1997), Digital acoustic analysis of precordial innocent versus ventricular septal defect murmurs in children. Am J Cardiol, 79(11):1552-55.
- Baura GD. (2005), The Business of innovation. *MX: The Online Information Source* for the Medical Device Industry.

- Bentley J. M., Grant P. M., J. McDonnel T. E. (1998), "Time- Frequency and Time-Scale Techniques for the Classification of Native and Bioprosthetic Heart Valve Sounds," *IEEE Trans. Biomedical. Eng.*, vol. 45, pp. 125-128., January 1998.
- Bishop, P.J. (1980). "Evolution of the Stethoscope", *Journal of the Royal Society of Medicine*.73:448- 456.
- Bonow R.O., Douglas L. M, Douglas P. Z. MD, Peter L. (2011), *Braunwald's Heart* Disease: Textbook of Cardiovascular Medicine. 9<sup>th</sup> ed. Saunders.
- Boric-Lubecke, O., Droitcour, A.D., Lubecke, V.M., Lin, J. and Kovacs, G.T.A. (2003) Wireless IC Doppler for sensing of heart and respiration activity. *IEEE Proceed- ings of International Conference on Telecommunication in Modern Satellite, Cable and Broadcasting Services*, Serbia and Montenegro, Nis, 1-3 October 2003, 337-344.
- Boutana, Daoud, M. Benidir, and B. Barkat (2014). "Segmentation And Time Frequency Analysis of Pathological Heart Sound Signals Using the EMD
   Method, *European Signal processing Conference*, Lisbon, Portugal.
- Brandstein M S, Ward E D B. (2001), *Microphone Arrays: Signal Processing Techniques and Applications*. Berlin: Springer-Verlag.
- Brusco M. and Nazeran H (2005), Development of an Intelligent PDA-based Wearable Digital Phonocardiograph, Proceedings of the 2005 IEEE Engineering in Medicine and Biology 27th Annual Conference, Shanghai, China, September 1-4.
- Bugtai N.T., Chan-Siy S.U., Chua J.E., Flores J.A., Wang J.L. (2012), Development of a Portable Heart Monitoring System, *IEEE TENCON Region 10 Conference*, Pp1-6, 19-22 Nov, Cebu, Philipina.
- Kwak C. O. Kwon W. (2012), Cardiac disorder classification by heart sound signals using murmur likelihood and hidden Markov model state likelihood, *IET Signal Process.*, Vol. 6, Iss. 4, pp. 326–334 doi: 10.1049/iet-spr.2011.0170
- Cable CS, Ducharme NG, Hackett RP, Erb HN, Mitchell LM, Soderholm LV (2002), Sound signature for identification and quantification of upper airway disease in horses. *Am J of Vet Res*, 63(12):1707-13.
- Cai, R., Lu, L., Hanjalic, A., Zhang, H.-J., and Cai, L.-H (2006). A flexible framework for key audio effects detection and auditory context inference. in *IEEE Trans. Speech Audio Processing*.

- Cai R., Lu L., Zhang H.-J. (2003), Using structure patterns of temporal and spectral feature in audio similarity measure, *Proc. ACM Multimedia*. 219-222.
- Cai, J.D., Yan, R.W.(2010), Application of FCM-HMM-SVM based mixed method for fault diagnosis of power electronic circuit. *Proc. Int. Conf. on Mechanic Automation and Control Engineering*, Wuhan, China, pp. 3982 – 3985
- Carter. G. (1977), Variance bounds for passively locating an acoustic source with a symmetric line array. *Journal of Acoustical Society of America*, 62(4): 922–926.
- Cheitlin, Alpert JS, Armstrong WF, Aurigemma GP, Beller GA, Bierman FZ, et al. (1997), ACC/ AHA guidelines for the clinical application of echocardiography: executive summary. A report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Committee on Clinical Application of Echocardiography). Developed in collaboration with the American Society of Echocardiography. *J Am Coll Cardiol*; 29:862-79.
- Chen Tian-hua Xing Su-xia Guo Pei-yuan Tang Hai-tao Yu Zheng (2010), The Design of Digital Collecting System of Heart Sound Signals Based on DSP, *International Conference on Biomedical Engineering and Computer Science* (*ICBECS*), 2010, pp. 1-4, Wuhan, China.
- Chen Tian-hua ,Xing Su-xia ,Guo Pei-yuan, (2011) "The Design of a New Digital Collecting System of Heart Sound Signals Based on XH-6 Sensor", International Conference on Measuring Technology and Mechatronics Automation.
- Christie, B.L (2009), Introduction to Biomedical Instrumentation. The technology of patient care. Cambridge University Press, Cambridge.
- Chung K, Zeng FG. (2009), Using hearing aid adaptive directional microphones to enhance cochlear implant performance, *Hearing Research*, 27-37.
- Chung K, Zeng FG., Acker K N. (2006), Effects of directional microphone and adaptive multichannel noise reduction algorithm on cochlear implant performance. *Journal of Acoustical Society of America*, 120: 2216-2227.
- Cohen L (2001): The uncertainty principle for the short time Fourier transform and wavelet transform. In *Wavelet transforms and time frequency signal analysis* Edited by Debnath L. Birkhäuser Boston, 1-428.
- Cohen L (1995), Time frequency analysis. Prentice Hall, Englewood Cliffs, NJ.

- Collins SP, Lindsell CJ, Peacock WF, Hedger VD, Askew J, Eckert DC, Storrow AB (2006), The combined utility of an S3 heart sound and B-type natriuretic peptide levels in emergency department patients with dyspnea. *J Card Fail*, 12(4):286-92.
- Cutnell, John, D. and Johnson, K.W. (1998) *Physics*. 4th Edition, John Wiley & Sons Ltd., New York.
- Barschdorff D., Femmer U., and Trowitzsch E. (1995), Automatic Phonocardiogram Signal Analysis in Infants Based on Wavelet Transforms and Artificial Neural Networks, Computers in Cardiology. Berlin, Germany: Springer-Verlag.
- De Bruijn NG (1973), A theory of generalized functions, with applications to Wigner distribution and Wyel correspondence. *Nieuw Archief voor Wiskunde*, XX1(3):205-80.
- DeGroff CG, Bhatikar S, Herzburg J, Shandas R, Vlades-Cruz L, Mahajan RL (2001), Artificial neural network-based method of screening heart murmurs in children. *Circulation*, 103:2711-2716.
- DeGroff,S, Bhatikar,J., Hertzberg,R, Shandas,L., Valdez-Cruz,and R. Mahajan (2001), Artificial neural network based method of screening heart murmurs in children, *Circulation*, pp. 2711–2716.
- Djebbari A., Bereksi-Reguig F. (2013), Detection of the valvular split within the second heart sound using the reassigned smoothed pseudo Wigner–Ville distribution, *Biomedical engineering online* 12 (1), 37
- Donnerstein R, Thomsen VV (1994), Hemodynamic and anatomic factors affecting the frequency content of Still's innocent murmur. *Am J Cardiol*, 74:508-510.
- Donghui, C., Zhijing, L.()2010, New text categorization method based on HMM and SVM. Proc. 2010 Second Int. Conf. on Computer Engineering and Technology, Chengdu, China, pp. 383 – 386
- Daoud B., Barkat. B and Benidir M. (2013), "Segmentation of pathological heart sound signal using empirical mode decomposition", *International journal of computer and electrical Engineering*, Vol.5, No 1.
- Daoud Boutana, M. Benidir and B. Barkat (2011), Segmentation and identification of some pathological phonocardiogram signals using time-frequency analysis, *Signal Processing, IET* Volume: 5, Issue: 6, Page(s): 527 -537.

- Digital Stethoscope Implementation on the TMS320C5515 DSP Medical SPRAB38A–June 2010, Development Kit (MDK).
- Dokur Z. and Olmez T. (2008), "ear sound classification using wavelet transform and incremental self-organizing map, Digital Signal Processing 18, Elsevier, pp. 951-959,
- Don Michael TA (1998): Auscultation of the Heart: A Cardiophonetic Approach. McGraw Hill, 392.
- Elfeky, M.G. Aref, W.G. Elmagarmid, A.K. (2005), WARP: time warping for periodicity detection. *In Fifth Intl. Conf. on Data Mining*, p. 8.
- Etchells E, Bell C, Robb K (1997), Does this patient have an abnormal systolic murmur? *JAMA*; 277:564-71.
- Flandrin P. and Martin W. (1997), Wigner-Ville spectrum of non-stationary random signals, *The Wigner Distribution*, Mecklenbrauker and Hlawatsch.
- Finley P, Warren AE, Sharratt GP, Amit M. (2006), Assessing children's heart sounds at a distance with digital recordings. *Pediatrics*, 118:2322-2325.
- Foote J., Matthew C. (2002). Audio Retrieval by Rhythmic Similarity. Proceedings of ISMIR, Paris, France, October 2002.
- Foote J. (1997), "Content-based retrieval of music and audio," *Proc of SPIE*, vol.3229, pp.138-147. (*ICASSP*), pp. 653-656.
- Guo Z., Moulder, C., Durand, L.-G. ; Loew, M. (1998), Development of a virtual instrument for data acquisition and analysis of the phonocardiogram, *Engineering in Medicine and Biology Society. Proceedings of the 20th Annual International Conference of the IEEE*, 436 439 vol.1
- Gupta C. N., Palaniapan R., Rajan S., Swaminathan S. (2007), "Neural network classification of homomorphic segmented heart sound", Applied Soft Computing 7, pp. 286-297.
- Gupta, C.N., Palaniappan, R., Swaminathan, S., Krishnan, S.M(2006), Neural network classification of homomorphic segmented heart sounds', *Appl. Soft Comput.*, 7, pp. 286 – 297
- Green JM, Wilcke JR, Abbott J, Rees LP (2006), Development and evaluation of methods for structured recording of heart murmur findings using SNOMED-CT post-coordination. J Am Med Inform Assoc, 13(3):321-33.
- Greenberg J E, Zurek p M. (1992), Evaluation of an adaptive beamforming method for hearing aids. *Journal of Acoustical Society of America*, 91: 1662-1676.

- Hedayioglu F., Jafari M. G., Mattos S.S., Plumbley M. D. and Coimbra M. T. (2012), Denoising and Segmentation of the Second Heart Sound Using Matching Pursuit, Proc. 34th Annual International Conference of the IEEE Engineering in Medicine and Biology Society (EMBC'12), San Diego, CA, USA, pp. 3440-3443, Doi 10.1109/EMBC.2012.6346705
- Hedayioglu F., Jafari M., Mattos S., Plumbley M., and Coimbra M. (2011), Separating sources from sequentially acquired mixtures of heart signals, *IEEE International Conference on Acoustics, Speech and Signal Processing* Hadrina Sh-Hussain, Sh- Hussain Salleh, A. Kamarul Ariff, Osama Alhamdani, Tan Tian-Swee, Alias Mohd Noor (2012), Application of Multipoint Auscultation for Heart Sound Diagnostic System (MAHDS), *The 11th International Conference on Information Sciences, Signal Processing and their Applications*.
- Hayek CS, Thompson WR (2003): Wavelet processing of systolic murmurs to assist with clinical diagnosis of heart disease. *Biomed Instrum Technol (Biomedical instrumentation & technology/Association for the Advancement of Medical Instrumentation)*, 37(4):263-70.
- Hebden, J.E., Torry, J.N. (1997), Identification of aortic stenosis and mitral regurgitation by heart sound analysis. *Computers in Cardiology*, 109–112.
- Henneberg S., Wiklund B.H, L.,and Sjodin G. (1992), Remote auscultatory patient monitoring during magnetic resonance imaging, *Journal of clinical monitoring*, vol. 8, pp. 37-43.
- Honghai Zhang, Maozhou Meng, Xiayun Shu, Sheng Liu (2006), "Design of a Flexible Stethoscope Sensor Skin Based on MEMS Technology", 7th International Conference on Electronics Packaging Technology,, Vol 23,pp 55-59.
- Huang Y T, Benesty J, Elko G W. (2000), Passive acoustic source localization for video camera steering. Proceedings of IEEE International Conference on Acoustic, Speech and Signal Processing, Istanbul, Turkey, 2: 909–912.
- http://www.littmann.com/wps/portal/3M/en\_US/3M-Littmann/stethoscope/littmannlearning-institute/auscultation-training/auscultation-techniques/
- Iacobellis, G., Willens, H.J., Barbaro, G., Sharma, A.M. (2008), Threshold values of high-risk echocardiographic epicardial fat thickness, *Obesity (Silver Spring)*, 160, pp. 887–892.

- Iead Rezek and Stephen J. Roberts (1998). Envelope Extraction via Complex Homomorphic Filtering. Research Report TR-98-9, June 1998.
- Ivashov, S.I., Razevig, V.V., Sheyko, A.P. and Vasilyev, I.A. (2004) Detection of human breathing and heartbeat by remote radar. *Progress in Electromagnetic Research Symposium*, Pisa, 28-31 March 2004, 663-666.
- Jia-Ren Chang Chien, Cheng-Chi Tai (2004), The Implementation of a Bluetooth-Based Wireless Phonocardio-Diagnosis System, *Proceedings IEEE International Conference on Networking, Sensing & Control*, pp 170-171.
- Jochen Moll, Christian Kexel, Viktor Krozer (2013), A Comparison of Beamforming Methods for Microwave Breast Cancer Detection in Homogeneous and Heterogeneous Tissue, *Proceedings of the 10th European Radar Conference*, Nuremberg, Germany.
- Jun Zhou, Yong He, Mohan Chirala, Brian M. Sadler, and Sebastian Hoyos (2013), Compressed Digital Beamformer With Asynchronous Sampling For Ultrasound Imaging, *IEEE International Conference on Acoustics, Speech* and Signal Processing (ICASSP), Vancouver, BC, Page: 1056 – 1060.
- K.J.ChoandH.H.Asada(2002), Wireless, battery-less stethoscope for wearable health monitoring (2002)," in IEEE 28th Annual Northeast Bioengineering Conference, April 20, 2002 - April 21, , Philadelphia, PA, United states, pp. 187-188.
- Kamarulafizam I.,Sh-Hussain Salleh,Rabia Bakhteri (2007), Heart Sound Analysis Using MFCC and Time Frequency Distribution," in *Proc. 3rd Intl. Conf. on Biomedical Eng.*, Kuala Lampur.
- Kaneda Y, Ohga J. (1986), Adaptive microphone-array system for noise-reduction. IEEE Transactions on Acoust, Speech and Signal Processing, 34(6): 1391-1400.
- Kassal J, Reeves W, Donnerstein R. (1994), Polymer-based adhered differential output sensor for cardiac auscultation. *Med Electronics*, 54-63.
- Ke Y., Hoiem D., and Sukthankar R. (2000). "Computer vision for music identification," in *Proc. CVPR*.
- Kim D, Tavel ME (2003), Assessment of severity of aortic stenosis through timefrequency analysis of murmur. *Chest*, 124:1638-1644.

- Klemm M., Craddock I. J., Leendertz J. A., Preece A., and R. Benjamin (2008), Improved Delay-and-Sum Beamforming Algorithm for Breast Cancer Detection, *International Journal of Antennas and Propagation*, Volume 2008, Article ID 761402,
- Klemm M., Leendertz J., Gibbins D., I. Craddock (2009), A. Preece, and R. Benjamin, "Microwave radar-based breast cancer detection: Imaging in inhomogeneous breast phantoms," *IEEE Antennas and Wireless Propagation Letters*, vol. 8, pp. 1349–1352.
- Kompis M, Dillier N. (2002), Performance of an adaptive beamforming noise reduction scheme for hearing aid application. *Journal of Acoustical Society of America*, 109: 1134-1143.
- Kovács F., Török M., Habermajer I. (2000), "A Rule-Based Phonocardiographic Method for Long-Term Fetal Heart Rate Monitoring," *IEEE Trans. Biomed. Engg.*, vol. 47., pp. 124- 130.
- Kudriavtsev VV, Kaelber D, Lazbin M, Polyshchuk VV, Roy DL (2006), New tool to identify Still's murmurs, 13th International Conference on Biomedical Engineering: ICBME.
- Lee, Y.M.; Moghavvemi, M (2002), "Remote heart rate monitoring system based on phonocardiography", *Student Conference on Research and Development*, Vol 78,pages:27–30,DOI: 10.1109/SCORED.2002.1033047.
- Leeuw A R, Dreschler W A. (1991), Advantages of directional hearing aid microphones related to room acoustics. Audiology, 330-344.
- Lembo NJ, Dell'Italia LJ, Crawford MH, O'Rourke RA (1988). Bedside diagnosis of systolic murmurs, *N Engl J Med.*, 318(24):1572-8.
- Liang H., Lukkarinen S., Hartimo I. (1997), Heart Sound Segmentation Algorithm Based on Heart Sound Envelogram, *Proc. Computers in Cardiology*, vol. 24.
- Lie Lu, Rui Cai, and Alan Hanjalic (2005). "Towards A Unified Framework for Content-based Audio Analysis", IEEE International Conference on Acoustics, Speech and Signal Processing (ICASSP) 2005, Vol. II, pp1069-1072.
- Liu L., Weilian Wang Gang Yuan, (2013) Algorithm of Heart Sound Segmentation and Feature Extraction, 2013 6th International Congress on Image and Signal Processing (CISP).

- Lu, G., Yang, F., Jing, X. and Wang, J. (2010) Contact- free measurement of heartbeat signal via a Doppler radar using adaptive filtering. *IEEE Proceeding of International Conference on Image Analysis and Signal Processing*, Hangzhou, 12-14 April 2010, 89-92.
- Lu L., Zhang H.-J., Jiang H. (2002), Content Analysis for Audio Classification and Segmentation, *IEEE Trans. on Speech and Audio Processing*, Vol.10, No.7, pp.504-516.
- Luis Torres-Pereira, Paulo Ruivo, Carla Torres-Pereira, Carlos Couto (1997), A Non-Invasive Telemetric Heart Rate Monitoring System Based On Phonocardiography, *IEEE ISIE*: 856-859.
- Mangione S, Nieman LZ. (1997), Cardiac auscultatory skills of internal medicine and family practice trainees. A comparison of diagnostic proficiency, *JAMA*., 278(9):717-22.
- Mansy H., Sandler R, Jones D (1999), "Testing Sensors for body surface vibration measurements", *Conference on Serving Humanity Advancing Technology*.
- Marcus GM, Gerber IL, McKeown BH, Vessey JC, Jordan MV, Huddleston M, McCulloch CE, Foster E, Chatterjee K, Michaels AD (2005), Association between phonocardiographic third and fourth heart sounds and objective measures of left ventricular function. JAMA, 293(18):2238-44.
- Matsumoto M, Hashimoto S (2006). A miniaturized adaptive microphone array under directional constraint utilizing aggregated microphones. *Journal of Acoustical Society of America*, , 119(1): 352-359.
- Matias Brusco, Homer Nazeran, Digital Phonocardiography (2004), A PDABased Approach, *Proceedings 26th Annual International Conference IEEE EMBS*, pp 2299-2302.
- McKee A. M. and Goubran R. A. (2005), Sound localization in the human thorax, Instrumentation and Measurement Technology Conference, Canada, vol. 1, pp.117–122.
- McKusick VA, Webb GN, Humphries JN, Reid JA (1955), On cardiovascular sound: further observations by means of spectral phonocardiography, *Circulation*, 11:849-870.
- McKusick VA (1958): Cardiovascular sound in health and disease. Baltimore: Williams & Wilkins.
- McKusick VA (1959), Spectral phonocardiography. Am J Cardiol, 4:200-206.

- McKusick VA, Massengale ON Jr, Wigod M, Webb GN (1956), Spectral phonocardiographic studies in congenital heart disease. *Brit Heart J*, 18:403-416.
- Messiah A. (1970), *Quantum mechanics*, Translated by G.M. Temmer, North Holland, Amsterdam, 1:50.
- Michael S Figueroa and Jay I Peters (2006), Congestive Heart Failure: Diagnosis, Pathophysiology, Therapy, and Implications for Respiratory Care, Respiratory Care, Vol. 51 No 4, pp 403-412.
- Mokhlessi, O., Rad, H.M., Mehrshad, N. (2010), Utilization of 4 types of artificial neural network on the diagnosis of valve-physiological heart disease from heart sounds. *Proc. 2010 17th Iranian Conf. on Biomedical Engineering*, pp. 1-4.
- Moll J. and Krozer V. (2012), On the optimization of ultra-wideband techniques for microwave breast cancer detection, in *International Sympo- sium on Signals*, *Systems and Electronics*, 2012, pp. 1–4.
- Moll J. and Krozer V. (2013), Time-Difference-of-Arrival Imaging for Ultra-Wideband Microwave Mammography, in 7th European Conference on Antennas and Propagation, Gothenburg, Sweden, pp. 2595–2599.
- Murray Longmore and Ian Wilkinson (2007), Oxford Handbook of Clinical Medicine (Oxford Medical Handbooks), Oxford University Press.
- Nakamitsu T., Shino H., Kotani T., Yana K., Harada K., Sudoh J., Harasawa E., and Itoh H. (1996), Detection and classification of systolic murmur for phonocardiogram screening, in *Proc. 18th Intl Conf. IEEE Eng. Med. Biol. Soc.*, pp. 123–124.
- Nelson P.B., Jin S.B., Carney A.E. (2003), Understanding speech in modulated interference: cochlaer implant users and normal-hearing listeners, J Acoust Soc Am., 113(2): 961-968.
- Nigam V, Priemer R (2006), A dynamic method to estimate the time split between the A2 and P2 components of the S2 heart sound. *Physiol Meas*, 27(7):553-67.
- Nigam V, Priemer R (2005), Accessing heart dynamics to estimate durations of heart sounds. *Physiol Meas*, 26(6):1005-18.

- Nigam V. and Priemer R. (2006),"A procedure to extract the aortic and the pulmonary sounds from the phonocardiogram", In Proceedings of the 28th IEEE EMBS Annual International Conference, pp. 5715-5718.
- Nichols M., Townsend N., Scarborough P., Rayner M., (2014), Cardiovascular disease in Europe: Epidemiological update, European heart journal.
- Noponen AL, Lukkarinen S, Angela A, Sikio K, Sepponen R (2000), How to recognize innocent vibratory murmur. *Comp in Cardiol*, 27:561-564.
- Noponen AL (2005), 4<sup>th</sup> Congress on Paediatric Cardiology. Poster Presentation.
- O'Halloran M., Glavin M., and E. Jones (2010), "Channel-ranked beamformer for the early detection of breast cancer," *Progress In Electromagnetics Research*, vol. 103, pp. 153–168.
- Obayya, M., Abou-Chadi, F.(2008), Classifying some cardiac abnormalities using heart rate variability signals. *Proc. 25th National Radio Science Conf.*, pp. 1-8.
- Padmanabhan V., Semmlow J.L.,and Welkowitz W. (1993) "Accelerometer type cardiac transducer for detection of low-level heart sounds," *IEEE Transactions on Biomedical Engineering, vol. 40*, pp.
- Pease, A.F.: If the heart could speak (2001),

w4.siemens.de/FuI/en/archiv/pof/heft2\_01/artikel19/index.html

- Phatiwuttipat P., Waree Kongprawechon, Kanokvate Tungpimolrut, Sumeth Yuenyong (2011), *The 8th Electrical Engineering Electronics, Computer, Telecommunications and Information Technology (ECTI).*
- Pipberger Hv, Stallman Fw (1962), Berson As. Automatic Analysis Of The P-Qrs-T Complex of The Electrocardiogram By Digital Computer. Ann Intern Med.; 57:776–787.
- Polyshchuk VV, Kudriavtsev VV (2005), Cardiovascular sound signature: method, process and format. USPTO Application, 2005; 0222515. Assignee: Biosignetics Corp; 1-41.
- Polyshchuk V, Choy FK, Braun MJ (2002), New gear-fault-detection parameter by use of joint time-frequency distribution. *AIAA J of Prop and Power*, 16(2):340-346.
- Polyshchuk V (1999): Detection and quantification of the gear tooth damage from the vibration and acoustic signatures, *PhD Dissertation*. University of Akron.

- Qian S, Dapang CD (1996), *Joint time-frequency analysis: method and applications*. Prentice Hall, NJ, 78-79.
- Rabiner L. and Shafer R. (1979), *Digital Processing of Speech Signals*, Prentice Hall: NJ.
- Rajan S., Doraiswami R., Stevenson R., and Watrous R. (1998), "Wavelet based bank of correlators approach for phonocardiogram signal classification," in *Proc. IEEE-SP Intl Symp. Time-Frequency and Time-Scale Analysis*, pp. 77– 80.
- Rakovi´c P., Sejdi´c E, Stankovi´c L. J., Jiang J (2006), Time-Frequency Signal Processing Approaches with Applications to Heart Sound Analysis, Computers in Cardiology; 33:197–200.
- Reed T., Reed N., and Fritzson P. (2001), The analysis of heart sounds for symptom detection and machine-aided diagnosis, *2nd Conf. Modeling and Simulation in Biology, Medicine, and Biomedical Engineering,* Delft, The Netherlands.
- Reed TR, Reed NE, Fritzson P (2004), Heart Sound analysis for symptom detection and computer-aided diagnosis. *Sim Mod Practice and Theory*, 12(2):129-46.
- Rein S, Reisslein M (2006), Identifying the classical music composition of an unknown performance with wavelet dispersion vector neural nets (extended version). *Information Sciences*, 176(12):1629-55.
- Rezek I., and Roberts S.J. (1998), *Envelope Extraction via Complex Homomorphic Filtering*. Research Report TR-98-9.
- Ritola J., Lukkarinen S. (1996), "Comparison of Time- Frequency Distributions in the Heart Sounds Analysis," *Medical & Biological Engineering & Computing*, vol 34., supplement 1., part 1., pp. 89-90., 1996.
- Sinha, R.K., Aggarwal, Y., Das, B.N. (2007), Backpropagation artificial neural network classifier to detect changes in heart sound due to mitral valve regurgitation, J. Med. Syst., pp. 205 – 209
- Saeid Saneia, Mansoureh Ghodsib, Hossein Hassanib (2011), An adaptive singular spectrum analysis approach to murmur detection from heart sounds, *Elsevier Journal of Medical Engineering & Physics 33*, pp. 362–367
- Say, O., Dokur, Z.; Olmez, T. (2002), Classification of heart sounds by using wavelet transform, 24th Annual Conference and the Annual Fall Meeting of the Biomedical Engineering Society EMBS/BMES, Volume 1, Issue, Page(s): 128 – 129.

- Schmidt, S.E., Toft, E., Holst-Hansen, C., Graff, C., Struijk, J.J.(2008), Segmentation of heart sound recordings from an electronic stethoscope by a duration dependent hidden-Markov model'. *Proc. Computers in Cardiology*, Bologna, Italy, pp. 345 – 348.
- Spiet A, Van Deun L, Eftaxiadis K, Laneau J, Moonen M, van Dijk B, van Wieringen A, Wouters J. (2007), Speech understanding in background noise with the two-microphone adaptive beamformer BETM in the Nucleus Freedom cochlear implant system. *Ear Hear*, 28(1): 62-72.
- Surr R K, Walden B E, Cord M T, Olsen L. (2002), Influence of environmental factors on hearing aid microphone preference. J. Am. Acad. Audiol., 13(6): 308-322.
- St Clair EW, Oddone EZ, Waugh RA, Corey GR, Feussner JR. Assessing housestaff diagnostic skills using a cardiology patient simulator. Ann Intern Med. 1992;117:751–756
- Tanya Chernyakova, Yonina C. Eldar, Ron Amit (2013), Fourier Domain Beamforming For Medical Ultrasound, IEEE International Conference on Acoustics, Speech and Signal Processing (ICASSP), Vancouver, BC, Page: 924-928
- Tavel ME. Cardiac auscultation (1996), A glorious past—but does it have a future? *Circulation* 1996;93:1250-3.
- Tavel ME, Katz H. (2005), Usefulness of a new sound spectral averaging technique to distinguish an innocent systolic murmur from that of aortic stenosis. Am J Cardiol, 95:902-904.
- Tavel ME (2006): Cardiac auscultation: a glorious past: and it does have a future. *Circulation*, 113:1255-1259.
- Tanveer Syeda-Mahmood, Fei Wang (2008), Shape-based retrieval of heart sounds for disease similarity detection, Computer Vision--ECCV, pp. 568--581, Springer Berlin Heidelberg.
- Thompson W., Hayek C., Tuchinda C., J. Telford, and J. Lombardo (2001), "Automated cardiac auscultation for detection of pathologic heart murmurs," *Pediatr. Cardiol.*, pp. 373–379.

- Tian-hua, C., Su-xia, X., Pei-yuan, G., Zheng, Y. (2009), The research of noninvasive method of coronary heart disease based on neural network and heart sound signal'. Proc. Int. Conf. on Information Engineering and Computer Science, Wuhan, China, pp. 1–4
- Tian Xian-ting, Zhao Zhi-dong (2011), Heart sound acquisition based on PDA and Bluetooth, *4th International Conference on Biomedical Engineering and Informatics (BMEI)*.
- Tilkian AG, Canover MB (1984), Understanding Heart Sounds and Mururs with an introduction to Lung Sound, Philadelphia, Sounders.
- Tovar-Corona B, Hind MD, Torry JN, Vincent R. (2001), Effects of respiration on heart sounds using time-frequency analysis. *Comp in Cardiol*, 457-460.
- Tovar-Corona B, Torry JN (1998), Time-frequency representation of systolic murmurs using wavelets. *Comput in Cardiol*, 601-604.
- Tuchinda C, Thompson WR (2001), Cardiac auscultatory recording database: delivering heart sounds through the internet. *Proc AMIA Symp*, 716-20.
- Vannuccini L., Earis J. E., Helistö P., Cheetham B. M. G., Rossi M., Sovijärvi A. R. A., and Vanderschoot J. (2000), Capturing and preprocessing of respiratory sounds, *Eur. Rsp Rev. 10*, Vol- 10, pp. 616-620.
- Várady P., Gross I., Hein A., Chouk L. (2001), Analysis of the Fetal Heart Activity by the Means of Phonocardiography, *Proc. IFAC Int. Conf. on Telematics and Automation*, Weingarten, Germany.
- Walden B E, Surr R K, Cord M T, Edwards B, Clson L. (2000), Comparison of benefits provided by different hearing aid technologies. J. Am. Acad. Aud., 11: 540-560.
- Wen-Zhu Wu et al (2009), Research on First Heart Sound and Second Heart Sound Amplitude variability and Reversal Phenomenon—A new finding in athletic heart study, Journal of Medical and Biological Engineering, 29 (4): 202-205.
- Wigner E (1932), On the quantum correction for thermodynamic equilibrium, *Phys Rev* 1932, 40:749-59.
- Wolfson H.J (1990), "On curve matching," in IEEE Trans. PAMI, pp.483-489, vol.12.
- Wood J. C., Barry D. T. (1995), "Time-Frequency Analysis of the First Heart Sound," *IEEE EMBS Magazine*, pp. 144-151., March-April 1995.

- Wouters J, Vanden Berghe J. (2001), Speech recognition in noise for cochlear implantees with a two-microphone monaural adaptive noise reduction system. Ear *Hear*, pp. 420-430.
- Xiao S., Cai S. and Liu G. (2000), "Studying the significance of cardiac contractility variability," IEEE Eng. Med. Biol. Mag., 19:102-105.
- Xiao, Y., Li, C. and Lin, J. (2007) A portable noncontact heartbeat and respiration monitoring system using 5-GHz radar. *IEEE Sensors Journal*, 7, 1042-1043. doi:10.1109/JSEN.2007.895979.
- Yi Zhang, Sixuan Chen, Jin Wu, Yuan Luo (2012), Remote Heart Sound and Lung Sound Monitoring System Design Based on ZigBee, 5th International Conference on BioMedical Engineering and Informatics (BMEI 2012), 16-18 October 2012, Chongqing, China.
- Yogeeswaran T. S., Daya D. D. N. B. and Wasudeva K. D. I. (2008), Construction of a Low Cost Electronic Heart Sound Monitoring System, *Proceedings of the Technical Sessions*, 72-77 Institute of Physics – Sri Lanka
- Yuanyang Li, Xinpei Wang and Churan Sun, Changchun Liu (2009), Identifying Heart Sound Sources through Multichannel, Acquisition and a 3-D Model, *Biomedical Engineering and Informatics*.
- Zeng F. G., Nie K., Stickney G.S., Kong Y.Y., Vongphoe M., Bhargave A., Wei C., Cao K. (2005), Speech recognition with amplitude and frequency modulations. Proc. Natl. Acad. Sci. USA, 102(7): 2293-2298.
- Zhenyu Guo, Chris Moulder, Louis-Gilles Durand, Murray Leow (1998),
  Development Of A Virtual Instrument For Data Acquisition And Analysis Of
  The Phonocardiogram, *Proceedings of the 20th Annual International Conference of the IEEE Engineering in Medicine and Biology Society*, Vol. 20, No 1.