Development of Breast Cancer Diagnosis Tool using Hybrid Magnetoacoustic Method and Artificial Neural Network

Maheza I. M. Salim, Abdul. H. Ahmad, Ismail Ariffin, Bustanur Rosidi, Eko Supriyanto.

Abstract-Breast cancer is a metabolic disease that causes the breast cells to acquire genetic alteration and allows them to grow beyond the normal tissue limit. With the yearly increasing trend in new cases and mortality rate, new approach in diagnosis and treatment of breast cancer is crucial to improve the existing management of breast cancer cases. This paper presents a new approach in breast cancer diagnosis by using Hybrid Magnetoacoustic Method (HMM) and artificial neural network. HMM is a newly developed one dimensional imaging system that combines the theory of acoustic and magnetism for breast imaging. It is capable to produce 2 outputs, the attenuation scale of ultrasound and the magnetoacoustic voltage. In this study, an artificial neural network is developed to automate the output of HMM for breast cancer classification. The ANN employs the steepest gradient descent with momentum back propagation algorithm with logsig and purelin transfer function. The best ANN architecture of 3-2-1 (3 network inputs, 2 neurons in the hidden layer, one network output) with learning rate of 0.3, iteration rate of 20000 and momentum constant of 0.3 was successfully developed with accuracy of 90.94% to testing data and 90% to validation data. The result shows the advantages of HMM outputs in providing a combination of bioelectric and acoustic information of tissue for a better breast cancer diagnosis consideration. The system's high percentage of accuracy shows that the output of HMM is very useful in assisting diagnosis. This additional capability is hoped to improve the existing breast oncology diagnosis.

Keywords—Breast Cancer, Hybrid Imaging, Artificial Neural Network

Manuscript received November 3, 2011: This work was supported in part by the Ministry of Higher Education of Malaysia and Universiti Teknologi Malaysia.

Maheza I. M. Salim is with the Faculty of Biomedical Engineering and Health Science, Universiti Teknologi Malaysia, 81310 Johor Bahru, Johor, Malaysia (+60197646095), (e-mail: <u>maheza@fke.utm.my</u>)

Ismail Ariffin and Abd. H. Ahmad are senior Lecturers at the Faculty of Electrical Engineering, Universiti Teknologi Malaysia, 81310 Johor Bahru, Johor, Malaysia (email: ismail@fke.utm.my, abdhamid@fke.utm.my)

Bustanur Rosidi is a Senior Lectrurer at the Institute of Bioproduct Development, Universiti Teknologi Malaysia, Johor Bahru, Johor, Malaysia (email: <u>bustanur@fke.utm.my</u>)

Eko Supriyanto is an Associate Professor at the Faculty of Biomedical Engineering and Health Science, Universiti Teknologi Malaysia, 81310 Johor Bahru, Johor, Malaysia. (eko@utm.my).

I. INTRODUCTION

A. Breast Cancer

Breast cancer is one of the most common cancers and the leading cause of cancer death among women worldwide [1, 4]. Breast cancer incidence is increasing over the years with more than 1 million new cases reported each year [3]. In addition to that, an average of 373000 women died globally every year in conjunction to the disease [2, 4]. In Malaysia specifically, National Cancer Registry report for the year 2003-2005 states that, the incidence rate of breast cancer in Malaysian population is 47.4 per 100000 populations with Chinese is at the highest rate of 59.9, followed by Indian at 54.2 and Malay at 34.9 per 100000 populations [10].

Breast cancer is a disease of uncontrolled breast cells growth, in which the cells acquire genetic alteration that allows them to multiply and grow outside the context of normal tissue development [36]. The cell metabolism increases to meet the requirements of rapid cell proliferation, autonomous cell growth and to maintain its survival [36].

The most common symptom of breast cancer is the presence of painless and slowly growth lump that may alter the contour or size of the breast [37-38]. It is also characterized by skin changes, inverted nipple and bloodstained nipple discharge [37-38]. The lymphatic nodes under the armpit may be swollen if affected by cancer. In late stage, the growth may ulcerate through the skin and infected [37-38]. Bone pain, tenderness over the liver, headaches, shortness of breath and chronic cough may be an indication of the cancer spreading to other organs in the body [37].

In the cancerous tissue, changes in density occur due to uncontrolled cell multiplication [36, 39-40, 42], excessive accumulation of protein in stroma [5-6, 41] and enhancement of capillary density [43, 44-46]. On the other hand, changes in conductivity also occur due to increase cellular water and electrolyte content as well as altered membrane permeability and blood perfusion to support metabolism requirements [5-6, 38, 41]. With the yearly increasing trend of breast cancer, improvement in diagnosis and treatment method is desirable to increase survival rate.

B. Ultrasonography in Breast Oncology Diagnostics

In of medical the world diagnostic. breast ultrasonography has an established and significant role in diagnostic of breast abnormalities [47]. Ultrasound is superior from mammography for its non-ionizing radiation. This makes ultrasound an imaging of choice to manage symptomatic breast in younger women as well as in pregnant and lactating mother whom the theoretical radiation of mammography is pertinent [11]. Ultrasonography is also a reliable modality for solid and cystic breast anomaly differentiation [12-13, 15, 47-49]. It is also used in imaging augmented and inflamed breast. However, in the current practice, the proportion of patient in whom breast ultrasonography is considered necessary is only 40% [15]. This means that, ultrasonography is not indicated for the rest 60% of patients referred for breast imaging [15]. This practice explains major constraint of ultrasonography in breast imaging that limits its usage for diagnostic of breast symptoms and for screening asymptomatic patients [11-15, 50-52].

The major problem of ultrasonography is its low sensitivity in detecting small and preinvasive breast cancers [11-15] from normal tissues due to the overlapping ultrasonic characteristics in these tissues [17-18, 33]. Breast ultrasound diagnostic relies on several sonographic features that are based on margin, shape and echotecture. Breast cancers are often characterized by poorly defined margins, irregular borders, spiculation, marked hyperechogenicity, shadowing and duct extension [11].

Another limitation of ultrasound is its inability to detect microcalcification, a calcium residue found in the breast tissue as an early indicator of certain type of breast cancer [12]. In ultrasonography, the presence of microcalcification in tissue is often masked with breast tissue heterogeneity and grainy noise due to speckle phenomena [53-54]. The reasons make microcalcification detection with ultrasonography unreliable [12].

In addition to that, previous study [52] reported that the sensitivity of ultrasonography for breast cancer detection is not only complicates by the low sensitivity of the ultrasound itself but also by the dependency of ultrasound result to operator. This means that, a single sonographic image may be interpreted differently by different operators and the result is relative to the operator skills and experience, variations in human perceptions of the images, differences in features used in diagnosis and lack of quantitative measures used for image analysis [55].

Due to the limitation of the existing ultrasound imaging system in breast cancer detection, an ultrasound-based hybrid system called the Hybrid Magnetoacoustic Method (HMM) has been developed in this study. The hybrid system was designed to be capable to access the acoustic and electric properties of tissue to get better diagnostic information.

C. Hybrid Magnetoacoustic Method

Biological tissue is a conductive element due to the

presence of random charges that support cell metabolism [57]. Propagation of ultrasound wave inside the breast tissue will cause the charges to move at high velocity due to the back and forth motion of the wave [55-60]. Moving charges in the present of magnetic field will experience Lorentz Force. Lorentz Force separates the positive and negative charges, producing an externally detectable voltage [55-57, 59] that can be collected using a couple of skin electrodes [55-57, 59].

Following the equation of wave propagation presented by Wen et al [63-65], the magnetoacoustic voltage amplitude is:

$V = WR_c B_0 \int_{Sound path final}^{Sound path final} \frac{\partial}{\partial z} \left[\frac{\sigma_z}{\rho_z}\right] \int_{-\infty}^t p(z, t) dt$ (1) [63-65]

in which, W is the ultrasound beam width, R_c is the resistance of the detection circuit , B_0 is the magnetic field intensity, σ is the samples conductivity, ρ is the samples density and p is the ultrasound pressure.

to the equation, the amplitude According of magnetoacoustic voltage is not directly proportional to the tissue conductivity, but is also weighted by the tissue density [63-65]. This is due to the fact that density is among the tissue parameters that determine the ultrasound attenuation and eventually reduce the velocity of ionic motion inside the tissue. For a heterogeneous tissue like breast cancer, the conductivity weighted density σ/ρ , is nonzero at tissue interface, giving rise to magnetoacoustic voltage. Hence, the magnetoacoustic voltage gives information that is relative to conductivity changes across interface [63-65] and this information can be used to access bioelectric properties of the breast tissue.

In addition to the magnetoacoustic voltage, HMM sensed back the ultrasound wave that is initially used to excite tissue ionic motion for tissue acoustic evaluation and gives information with regards to the acoustic attenuation scale of the breast tissue.

Therefore, HMM is designed to be capable to access the conductivity and density of breast tissues. The block diagram of HMM is shown by figure 1.1.



Fig. 1.1: Block diagram of HMM system. (a) Side view, (b) Cross sectional view

A series of experiment and quantification on the output of HMM to 24 normal and 25 cancerous mice breast tissue show that the combination of acoustic and bioelectric properties is a promising way of breast cancer diagnostic. The result shows that normal mice breast tissues experience the highest attenuation $(2.329\pm1.103 \text{ dBmm}^{-1})$, followed by cancerous tissue $(1.76\pm1.08 \text{ dBmm}^{-1})$ with the difference of 0.569 ± 0.023 dB. In addition to that, mean magnetoacoustic voltage results for tissue the normal and cancerous tissue group are $0.42\pm0.16 \mu$ V and $0.8\pm0.21 \mu$ V respectively.

Therefore, this expansion of study on the development of breast cancer diagnosis system by using hybrid magnetoacoustic method and artificial neural network will concentrate on developing an automated system for HMM outputs based on the experimental result.

D. Artificial Neural Network

Artificial intelligence has been used very extensively in modeling biomedical application. It has been proposed as reasoning tool to support clinical decision making since the earliest day of computing. An artificial neural network (ANN) is a nonlinear and complex computational mathematical model for information processing with architectures inspired by neuronal organizational biology [19-21]. The underlying reason for using an artificial neural network in preference to other likely methods of solution is its ability to provide a rapid solution. Depending on the type of problem being considered, ANN is a proven method which is a capable of providing fast assessment and accurate result [19-21]. This is because; ANN works in a simulated parallel manner and is not limited by the serial requirements of the normal program such as in expert system and conventional programming [19-22].

The most valuable property of ANN is its ability to learn and to generalize [22]. Generalization refers to the capability of neural network to produce reasonable outputs for input which is not encountered during training [19, 23]. These attributes mark neural network out from other computational methods [22]. Neural network consists of a number of simple and highly connected processors. Like the brain, ANN can recognize pattern, manage data and most significantly, learn [19-20]. Previous study also showed that, ANN with at least one hidden layer of computational unit is capable of approximating any finite function to any degree of accuracy as a universal approximator [24].

In medicine, ANN is widely used for modelling, data analysis and diagnostic classifications [19-21, 24-25]. The most common ANN model used in clinical medicine is the multilayer perceptron (MLP) [25]. The most widely used connection pattern is three layer back propagation neural network which have been proved to be useful in modelling input –output relationship [69-70, 77] while the most commonly used transfer functions are linear, log sigmoid and tan sigmoid [78].

The most commonly used indicator of ANN performance is Mean Squared Error (MSE). MSE is the average of the squares of the difference between each output and the desired output. Research performed in [19-20, 23, 25-26, 28, 61] used MSE as the measurement of ANN performance. In addition to that, researches conducted in [23, 61, 29-31] were using the accuracy of the tested data as one of the performance indicator of the ANN. By using this method, network is trained using a part of the data and the remainder is assigned as the testing and validation data.

II. METHODOLOGY

The HMM system is capable to produce 2 outputs: The attenuation scale of ultrasound and the magnetoacoustic voltage. In this study, ultrasound attenuation scale and magnetoacoustic voltage data from 24 normal and 25 cancerous mice breast tissue samples were used. The breast tissues were harvested from a set of tumor bearing mice FVB/N-Tg MMTV PyVT 634 Mul that carries invasive adenocarcinoma and its control strain FVB/N that carries normal breast tissue.

A. Ultrasound Attenuation Scale

The ultrasound attenuation scale is the degree of weakening of ultrasound amplitude and intensity as it propagates through a medium. In HMM, the ultrasound attenuation scale is determined following the insertion loss method. During the ultrasound measurement, vegetable oil was used as medium for ultrasound propagation and the tissue was immersed into the oil that is located between the ultrasound transmitter and receiver. To measure the attenuation scale of ultrasound in tissue, the power spectral density (PSD) of ultrasound wave propagating through the oil medium was first calculated, followed by the PSD of ultrasound propagating through the oil and tissue. The final attenuation was determined by calculating the difference of PSD between the oil medium and oil medium with tissue at 9.8MHz. The total number of attenuation scale used in the development of breast cancer diagnosis system is presented by Table 2.1

No	Attenuation scale	Quantity
1	Cancerous tissue	106
2	Normal tissue	106

Table 2.1: Number of ultrasound attenuation scale at 9.8MHz

B. Magnetoacoustic Voltage Data

The magnetoacoustic voltage data is the voltage that results from Lorentz Force interaction to conductive charges in tissue. Its amplitude is influenced by tissue conductivity and tissue attenuation level. In HMM, magnetoacoutic voltage measurement was made by touching the tissue surface from the x direction by using the skin electrodes. The total number of magnetoacoustic voltage signals that were recorded in this experiment is presented by Table 2.2.

No	Magnetoacoustic voltage	Quantity	
	signal		
1	Magnetoacoustic voltage of Normal Breast Tissue (1&2)	212	
2	Magnetoacoustic voltage of Cancerous Breast Tissue (1&2)	212	

Table 2.2: Total number of magnetoacoustic voltage signal recorded by HMM.

C. Data Massaging

Data massaging involves restructuring the range of neural network input and target values between zero to one. Massaging is done due to the fact that neural network works best when all its input and output vary within the range of 0-1 [19-20].

D. Data Sampling

Data comprises of 25 cancerous tissues and 24 normal tissues were arranged randomly into the training, testing and

validation data shown by Table 2.3. For the ultrasound data, measurement was repeated 5 times for every sample. On the other hand, for magnetoacoustic voltage data, measurement was repeated 10 times.

ANN Database	Ultrasound data		Magnetoacoustic voltage data			
		Side		e 1	Side 2	
	N	С	N	C	Ν	C
Training	68	69	68	69	68	69
Testing	28	27	28	27	28	27
Validation	10	10	10	10	10	10
Total	106	106	106	106	106	106

*N- Normal, C-Cancerous

Table 2.3: ANN data sampling

E. Development of Artificial Neural Network

The diagnosis of breast cancer in this study was performed by employing a Multilayer Feed Forward Neural Network (MFNN) with 2 inputs. It was trained by using the *steepest descent with momentum back propagation* algorithm with *logsig* and *purelin* transfer function in Matlab environment. The back propagation algorithm is the most commonly used algorithm in medical computational application as were experimented by [19-20, 25, 29].

The measurement of ANN performance was observed by using the Mean Squared Error (MSE) and total classification accuracy of the network to the testing data. Training is best when the ANN is capable to achieve lowest MSE value.

In addition to that, each ANN configuration was tested by using the testing group data to obtain the overall classification accuracy as were experimented in [19-20, 29-30]. By using this method, network was trained using a part of the data and the remainder was assigned as testing and validation data.

The overall flow chart for the development of MFNN in this study is shown by figure 2.1.



Fig 2.1: Overall flow chart for the development of breast cancer diagnosis tool in this study.

III. RESULT

A. Results for the determination of optimized number of neuron in hidden layer.

During the hidden neuron optimization step, the number of neuron in the hidden layer was varied from 1 to 12 while the other parameters including learning rate, iteration rate and momentum constant were fixed to a predetermined value of 0.3, 20000 and 0.2 respectively. From figure 3.1, it can be observed that the lowest MSE value that was achieved during the neuron optimization was given by network with 12 hidden neurons at 0.098. However, its classification accuracy is only 89.09%. On the other hand, the highest classification accuracy of 90.94% was given by 2 hidden neurons with MSE of 0.111. An increase of 0.013 in MSE has given 2% increments in classifications accuracy. Hence, hidden layer size of 2 was chosen for optimum number of neuron in the hidden layer for the ANN with slight compensation in higher MSE value.



Figure 3.1: Number of neuron in the hidden layer vs Mean Squared Error (MSE) and the total accuracy (%)

B. Result for the determination of optimized learning rate

The ANN learning rate was varied from 0.1 to 0.9 during the learning rate optimization step. Training epochs and momentum constant were kept at their predetermined value of 20000 and 0.2 respectively. The number of neuron in the hidden layer was set to its optimum value of 2 neurons. Figure 3.2 shows that the lowest MSE value was achieved with learning rate of 0.1 to 0.8. In addition to that, learning rate of 0.3 to 0.8 gives the highest classification accuracy of 90.94%. In this case, 6 optimum values of learning rate were obtained with the same MSE value and classification accuracy to testing data. Hence, the best learning rate was chosen by testing the robustness of the ANN to validation data. The validation result shows that learning rate of 0.3 gives 90% accuracy compared to learning rate of 0.4 to 0.8 that gives 85% validation accuracy. Hence, learning rate of 0.3 was chosen as the best learning rate values.



Figure 3.2: Learning rate vs Mean Squared Error (MSE) and the total accuracy (%)

C. Result for the determination of optimized iteration rate

The iteration rate was varied from 5000 to 50000 with a constant increment of 5000 during the learning rate optimization process. Other parameters were set at their optimum value except momentum constant, which was at its predetermined value of 0.2. Figure 3.3 shows that iteration rate which is too short produces ANN with high MSE and low classification accuracy. This indicates that the iteration rate is insufficient to allow the network to converge. It also shows that iteration rate of 20000, 40000 and 50000 give the lowest MSE value of 0.111 and the highest classification accuracy of 90.94%. Hence, iteration rate of 20000 was chosen since this architecture produces lowest MSE and highest classification accuracy at the shortest time interval. In addition to that, this architecture was tested to validation data and it gives the highest validation classification accuracy of 90%.



Figure 3.3: Iteration rate vs Mean Squared Error (MSE) and the total performance accuracy (%)

C. Result on the determination of optimized momentum constant

The final step for the determination of optimum ANN was to find the best momentum constant for the network. The momentum constant was varied from 0.1 to 0.9 while other parameters were kept at their optimum values. Figure 3.4 indicates that momentum constant of 0.2, 0.3 and 0.5 produces network with lowest MSE value. Among that, momentum constant of value 0.3 and 0.5 gives the highest classification accuracy on testing data. Hence, validation data was used to test the robustness of the system and the result indicates that momentum constant of 0.3 gives highest validation classification accuracy of 90%.



Figure 3.4: Momentum constant vs Mean Squared Error (MSE) and total performance accuracy (%).

The final architecture of ANN for classifications of breast cancer in this study is 3-2-1 (3 network inputs, 2 neurons in the hidden layer, one network output) with learning rate of 0.3, iteration rate 0f 20000 and momentum constant of 0.3.

The final classification performance of the optimum ANN for testing and validation data is shown in Table 3.1. The result indicates that the ANN is capable to achieve 90.94% and 90% classification accuracy for testing and validation data. This result shows the advantages of HMM output in providing additional bioelectric parameter of tissue for breast cancer diagnosis consideration. The system's high percentage of accuracy shows that the output of HMM is very useful in assisting diagnosis. This additional capability is hoped to improve the existing breast oncology diagnosis.

Data	Testing Data		Validation Data	
	Normal	Cancer	Normal	Cancer
Actual Data	28	27	10	10
ANN Result	25	25	8	10
% Accuracy	89.29	92.59	80	100
% Total Accuracy	90.94		90	

Table 3.1: Classification Result of the Neural Network

IV. CONCLUSION

ANN with architecture of 3-2-1 (3 network inputs, 2 neurons in the hidden layer, one network output) with learning rate of 0.3, iteration rate of 20000 and momentum constant of 0.3 was successfully developed with accuracy of 90.94% to testing data and 90% to validation data. The result shows the advantages of HMM outputs in providing a combination of bioelectric and acoustic information of tissue for breast cancer diagnosis consideration.

References:

- J. Ferlay, F. Bray, P. Pisani, D. M. Parkin, "Cancer incidence, mortality and prevalence worldwide", *IARC CancerBase*, vol. 5(2), 2004.
- [2] D. M. Parkin, F. I. Bray, S. S. Devessa, "Cancer burden in the year 2000. The global picture", *Eur J Cancer*, vol 37, pp 54-66, 2001.
- [3] B. McAree, M. E. O'Donnell, A. Spence, T. F. Lioe, D. T. McManus, R. A. J. Spence, "Breast Cancer in Women under 40 Years of Age: A Series of 57 Cases from Northern Ireland", *The Breast*, vol 19. Pp. 97-104, 2010.
- [4] V. Naraynsingh, S. Hariharan, D. Dan, S. Bhola, K. Nagee, "Trends in breast cancer mortality in Trinidad and Tobago—A 35-year study", *Cancer Epidemiology*, vol. 34, pp. 20-23, 2010.
- [5] M. J. Bissel, D. Radisky, "Putting tumors in context. Nat Rev Cancer, vol. 1, pp. 46-54, 2001.
- [6] E. Cukierman, "A visual quantitative analysis of fibroblastic stromagenesis in breast cancer progression", J Mammary Gland Biol Neoplasia, vol 9, pp. 311-324, 2004.
- [7] X. Sim, A. Ali, S. Wedren, D. L. M. Goh, C. S. Tan, M. Reilly, "Ethnic differences in time trend of female breast cancer incidence: Singapore", *BMC Cancer*, vol. 6, 2006.
- [8] V. M. Medina, A. Laudico, M. R. Mirasol-Lumague, H. Brenner, M. T. Redaniel, "Cumulative incidence trends of selected cancer sites in a Philippine population from 1983 to 2002: a joinpoint analysis". *Br J Cancer*, vol 102, 2010.
- [9] J. M. Park, "Promising Techniques for Breast Cancer Detection, Diagnosis and Staging Using Non-Ionizing Radiation Imaging Techniques", *Physica Medica*, vol XXI, supplement 1, 2006.
- [10] RadiologyMalaysia, http://www.radiologymalaysia.org, 8th June 2011.
- [11] W. The, A. R. M. Wilson, "The Role of Breast Ultrasound in Breast Cancer Screening. A Consensus Statement by The European Group for Breast Cancer Screening", *European Journal of Cancer*, vol. 34(4), pp. 449-450, 1998.
- [12] C. M. Sehgal, S. P. Weinstein, P. H. Arger, E. F. Conant, "A Review of Breast Ultrasound", *J Mammary Gland Biol Neoplasia*, vol 11, pp. 113-123, 2006.
- [13] L. W. Bassett, C. Kimme-Smith, L. K. Sutherland, R. H. Gold, D. Sarti, W. King, "3rd Automated and Hand Held Breast Ultrasound: Effect on Patient Management", *Radiology*, vol. 165, pp. 103-108, 1987, 165.
- [14] P. H. Arger, C. M. Sehgal, E. F. Conant, J. Zuckerman, S. E. Rowling, J. A. Patton, "Inter Reader Variability and Predictive Value of US Descriptions of Solid Breast Masses: Pilot Study", *Acad Radiol*, vol. 8, pp. 335-342, 2001.
- [15] K. Flobbe, P. J. Nelemans, A. G. H. Kessels, G. L. Beets, M. F. Von Meyenfeldt, J. M. A. Van Engelshoven, "The Roll of Ultrasonography as an Adjunct to Mammography in the Detection of Breast Cancer: A Systematic Review", *European Journal of Cancer*, vol. 38, pp. 1044-1052, 2002.
- [16] J. C. Bamber, "Ultrasonic Propagation Properties of the breast", in Ultrasonic Examination of the breast, John Wiley and Sons, 1983.
- [17] P. D. Edmonds, and C. L. Mortensen, "Ultrasonic Tissue Characterization for Breast Biopsy Specimen", *IEEE Ultrasonic Symposium*, pp. 915, 1987.
- [18] L. Landini and S. Sarnelli, "Evaluation of the Attenuation Coefficient in Normal and Pathological Breast Tissue", *Medical and Biological Engineering and Computing*, vol. 24, pp. 243-247, 1986.

- [19] F. Ibrahim, "Prognosis of Dengue Fever and Dengue Hemorrhagic Fever Using bioelectrical Impedance", Ph.D Thesis, Department of Biomedical Engineering, Faculty of Engineering, University of Malaya; 2005.
- [20] F. Ibrahim, T. Faisal, M. I. Mohamad Salim, M. N. Taib, "Non Invasive Diagnosis of Risk in Dengue Patients Using Bioelectrical Impedance Analysis and Artificial Neural Network", *Med Biol Eng Compute*, vol. 48, pp. 1141-1148, 2010.
- [21] D. E. U beyli, "Implementing Automated Diagnostic System for Breast Cancer Detection", *Expert System with Application*, vol. 33, pp. 1054-1062, 2007.
- [22] Introduction to Artificial Neural Network, IEEE 1995, 08186 7085 -1/95.
- [23] S. Haykin, "Neural Networks: A Comprehensive Foundation", New York; McMillan, 1994.
- [24] M. Sun, R. J. Sclabassi, "The Forward EEG Solutions can be computed using Artificial Neural Networks", *IEEE Transactions on Biomedical Engineering*, vol. 47, pp. 1044-1050, 2000.
- [25] T. John, Z. Z. Wei, S. D. Barnhill, R. Madyastha, "Understanding Artificial Neural Networks and Exploring their Potential Applications for the Practicing Urologist', *Urology*, vol. 55, pp. 161-172, 1998.
- [26] E. Haselsteiner, G. Pfurtscheller, "Using Time-Dependent Neural Networks for EEG Classification", *IEEE Trans Rehabil Eng*, vol. 8, pp. 457-463, 2000.
- [27] M. Negnevitsky, "Artificial Intelligence, A Guide to Intelligent System", Pearson Education, 2002. ISBN: 0201711591
- [28] S. K. Sinha and P. W. Fieguth, "Projection Neural Network Model for Classification of Pipe Defects", *J Autom Constr*, vol. 15(1), p. 73, 2005.
- [29] S. Y. Joo, W. K. Moon, H. C. Kim, "Computer Aided Diagnosis of Solid Breast Nodules on Ultrasound with Digital Image Processing and Artificial Neural Network", IEEE, 2004, 0 - 7803 - 8439 - 3/04
- [30] R. I. Lammers, D. L. Hudson, M. E. Seaman, "Prediction of Traumatic Wound Infection with a Neural Network-Derived Decision Model", *American Journal of Emergency Medicine*, vol 21, 2003.
- [31] A. Subasi, "Automatic Recognition of Alertness Level from EEG by using Artificial Neural Network and Wavelet Coefficient", *Expert* System with Applications, vol. 28, pp. 701-711, 2005.
- [32] M. E. Anderson, M. S. C. Soo, G. E. Trahey, "Optimizing Visualization for Breast Microcalcifications", *IEEE Ultrasonic Symposium*, pp. 1315-1320, 2000.
- [33] M. I. Mohamad Salim, S. N. Z. Ahmmad, B. Rosidi, I. Ariffin, A. H. Ahmad, E. Supriyanto, "Measurements of Ultrasound Attenuation for normal and pathological mice breast tissue Using 10MHz Ultrasound Wave", *Proceeding of The 3rd WSEAS International Conference on Visualization, Imaging and Simulation* (VIS'10), 2010.
- [34] M. I. Mohamad Salim, M. A. Tumiran, S. N. Makhtar, B. Rosidi, I. Ariffin, A. H. Ahmad, E. Supriyanto, "Hybrid Magnetoacoustic Method for Breast Tumor Detection: An In-Vitro and In-vivo Modelling and Analysis". WSEAS Transactions on Information Science and Applications, vol. 7(8), pp. 1048-1057, 2010.
- [35] M. I. Mohamad Salim, M. A. Tumiran, S. N. Makhtar, B. Rosidi, I. Ariffin, A. H. Ahmad, E. Supriyanto, "Quantitative analysis of Hybrid Magnetoacoustic Method for Detection of normal and pathological Breast Tissue", *Proceeding of 12th WSEAS International Conference on Automatic Control, Modelling and Simulation*, pg 144-149, 2010.
- [36] J. W. Locasale, and L. C. Cantley, "Altered Metabolism in Cancer", BMC Biology, vol. 88, pp. 88, 2010.
- [37] Breast cancer, Malaysia Oncology Society, malaysianoncology.org, 21st May 2011.
- [38] Y. Zou. Z. Guo, "A Review of Electrical Impedance Techniques for Breast Cancer Detection", *Medical engineering and physics*, vol. 25, pp. 79-90, 2003.
- [39] J. Jossinet, A. Lobel, C. Michoudet, M. Schmitt, "Quantitative Technique for Bio-Electrical Spectroscopy", *J Biomed Eng*, vol. 7, pp. 289-294, 1985.
- [40] J. Jossinet, "Variability of Impedivity in Normal and Pathological Breast Tissue", Med Biol Eng Comput, vol. 34, pp. 346-350, 1996.
- [41] M. L. Arendt, J. A. Rudnick, P. J. Keller, C. Kuperwasser, "Stroma in Breast Development and Disease", Seminar in Cell and Developmental

Biology, vol. 21, pp. 11-18, 2010.

- [42] C. Tophkhane, S. Yang, Z. J. Zhao, X. Yang, "Cell density-dependant regulation of p73 in breast cancer cells", *International Journal of Breast Oncology*, vol. 35, pp. 1429-1434, 2009.
- [43] J. K. Chan, A. Magistris, V. Loizzi, F. Lin, J. Rutgers, K. Osann, P. J. DiSaia, M. Samoszuk, "Mast cell density, angiogenesis, blood clotting, and prognosis in women with advanced ovarian cancer", *Gynecologic Oncology*, vol. 99, pp. 20-25, 2005.
- [44] M. B. Donati, A. Falanga, "Pathogenetic mechanism of thrombosis in malignancy", Acta Haematol, vol. 106(1-2), pp. 18-24, 2001.
- [45] F. R. Rickles, A. Falanga, "Molecular Basis for the relationship between thrombosis and cancer", *Thromb Res*, vol. 102(6), pp. 215-224, 2001.
- [46] M. Shinoji, W. W. Hancock, K. Abe, C. Micko, K. A. Casper, R. M. Baine, "Activation of coagulation and angiogenesis in cancer: immunohistochemical localization in situ of clotting proteins and vascular endothelial growth factor in human cancer", *Am J Pathol*, vol 152(2), pp. 399-411, 1998.
- [47] W. E. Svensson, "A Review of the Current Status of Breast Ultrasound", European Journal of Ultrasound, vol. 6, pp. 77-101, 1997.
- [48] H. Laine, J. Rainio, H. Arko, T. Tukeva, "Comparison of Breast Structure and Findings by X- Ray Mammography, Ultrasound, Cytology and Histology: a Retrospective Study", *Eur J Ultrasound*, vol 2, pp. 107-115, 1995.
- [49] A. T. Stavros, D. Thickman, C. L. Rapp, M. A. Dennis, S. H. Parker, G. A. Sisney, "Solid Breast Nodules: Use of Sonography to Distinguish between Benign and Malignant Lesions", *Radiology*, vol. 196, pp. 132-134, 1995.
- [50] K. Singh, T. Azad, G. Dev Gupta, The Accuracy of Ultrasound in Diagnosis of Palpable Breast Lump, *JK Science*, vol. 10(4), pp. 186-188, 2008.
- [51] A. T. Stavros, "Breast Ultrasound", Philadelphia: William and Wilkins, 2004.
- [52] J. M. Chang W. K. Moon, N. Cho, J. S. Park, S. J. Kim, "Radiologist's Performance in the Detection of Benign and Malignant Masses with 3D Automated Breast Ultrasound (ABUS)", *European Journal of Radiology*, vol. 78, pp. 99-103, 2011.
- [53] S. P. Weinstein, C. M. Seghal, E. F. Conant, J. A. Patton, "Microcalcifications in breast tissue phantoms visualized with acoustic resonance coupled with power Doppler US: initial observations", *Radiology*, vol. 224, pp. 265-269, 2002.
- [54] A. Alizad, M. Fatemi, L. E. Wold, J. F. Greenleaf, "Performance of vibro-acoustography in detecting microcalcifications in excised human breast tissue: a study of 74 tissue samples", *IEEE Trans Med Imag*, vol. 23, 307-312, 2004.
- [55] H. Wen, J. Shah, R. S. Balaban, "Hall Effect Imaging", IEEE Transactions on Biomedical Engineering, vol. 45, pp. 119-124, 1998.
- [56] H. Wen, E. Bennett, J. Shah, R. S. Balaban, "An Imaging Method using Ultrasound and Magnetic Field" *IEEE Ultrasonic Symposium*, pp. 1407, 1997.
- [57] H. Wen, E. Bennet, "The Feasibility of Hall Effect Imaging In Humans" IEEE Ultrasonic Symposium, pp. 1619, 2000.
- [58] A. Montalibet, J. Jossinet, A. Matias, D. Cathignol, "Interaction Ultrasound –Magnetic field: Experimental Set Up and Detection of the Interaction Current", *IEEE Ultrasonic Symposium*, pp. 533, 2000.
- [59] Y. Su, S. Haider, A. Hrbek, "Magnetoacousto Electrical Tomography, A new Imaging Modality for Electrical Impedance", *IFMBE Proceeding* vol. 17, pp. 292, 2007.
- [60] S. J. Norton. "Can Ultrasound be used to Stimulate Nerve Tissue?" Biomedical Engineering Online, vol 2, pp. 1-9, 2003.
- [61] I. A. Basheer, M. Hajmeer, "Artificial Neural Networks: Fundamentals, Computing, Design, and Application", *J Microbiol Methods*, vol. 43, pp. 23-31, 2000.

Maheza I. M. Salim was born in Selangor, Malaysia, on 26th August 1983. She received an honour degree in Biomedical Engineering (with Distinction) from The University of Malaya, Malaysia in 2006. Currently, she is pursuing her Ph.D degree in Medical Imaging in Universiti Teknologi Malaysia with specialization in hybrid imaging modality for breast cancer detection.

Ismail Ariffin was born in Penang, Malaysia, on the 13th September 1962. He received a Bachelor of Science degree in Electronic and Electrical Engineering from University of Miami, USA in 1987 and a master degree in Electrical Engineering (Specialization in Electronic) from Universiti Teknologi Malaysia in 1991. Currently he is a senior lecturer at the Faculty of Electrical Engineering, Universiti Teknologi Malaysia. He joined the Faculty of Electrical Engineering at Universiti Teknologi Malaysia since 1983. He is currently a Malaysian Standard (SIRIM) technical committee on Anaesthetic/Respiratory and Electronic, medical imaging, digital signal processing and image processing.

Abd Hamid Ahmad was born in Melaka, Malaysia, on the 14th July 1960. He received an honours degree in Electrical and Electronic Engineering in1986 from University of Miami and a postgraduate honours degree in Electronic in1991, from Universiti Teknologi Malaysia. He joined the Electronics Department, Faculty of Electrical Engineering, Universiti Teknologi Malaysia, as Assistant Lecturer in 1987, teaching Electronics, Digital Electronics, Digital System, Microprocessor and Electronic System. His current research interests include digital system design, electronics system design and computer assisted learning.

Bustanur Rosidi was born in Nganjuk, Indonesia on 9th February 1966. He received his Diplom-Biochemiker (degree and MSc in biochemistry) at University of Hannover, Germany in 1994. Between 1995 and 1997 he worked as researcher on a project sponsored by DFG at Department of Molecular Neuro Biochemistry, Ruhr-University of Bochum, Germany. He pursued his PhD at Department of Molecular Genetics University of Hannover between 1997 and 2001, where he graduated in July 2001. His PhD work focused on immune biochemical and molecular biological characterization of antigens originating from plants. In 2002 he joined the group of G. Iliakis at Institute of Medical Radiation Biology, University Medical School, University of Duisburg-Essen, in Essen Germany, as a post doctoral researcher working on the biochemistry of DNA repair, especially on the mechanism of non-homologous endjoing (NHEJ). In November 2007 he joined the University Technology Malaysia (UTM) in Johor Bahru, Malaysia, where he is currently working as lecturer at the Institute of Bioproduct Development (IBD).

Prof. Eko Supriyanto is a biomedical engineer. He is head of Diagnostics Research Group, Universiti Teknologi Malaysia. He obtained his PhD in medical electronics from University of Federal Armed Forces, Hamburg, Germany. His research interest is engineering application in medicine. He has more than 130 international publications in the area of medical electronics, medical image processing and medical computing. He has 22 national and international innovation awards and more than 10 patents of biomedical products.