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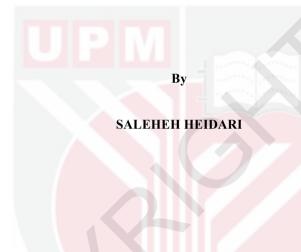
REGIONAL CONTRAST ENHANCEMENT AND FOUR-DIRECTIONAL THRESHOLDING TECHNIQUES FOR PULMONARY NODULE EXTRACTION AND DISCRIMINATION

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FSKTM 2015 10



REGIONAL CONTRAST ENHANCEMENT AND FOUR-DIRECTIONAL THRESHOLDING TECHNIQUES FOR PULMONARY NODULE EXTRACTION AND DISCRIMINATION



Thesis Submitted to the School of Graduate Studies, Universiti Putra Malaysia, in Fulfillment of the Requirements for the Degree of Master of Science

July 2015

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A special dedication to my loving family

Abstract of thesis presented to the Senate of Universiti Putra Malaysia in fulfilment of the requirement for the degree of Master of Science

REGIONAL CONTRAST ENHANCEMENT AND FOUR DIRECTIONAL THRESHOLDING TECHNIQUES FOR PULMONARY NODULE EXTRACTION AND DISCRIMINATION

By

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July 2015

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Automated pulmonary nodules extraction and lung disease diagnosis by Computer Aided Diagnosis (CAD) systems is a challenging task. Generally, the CAD system utilizes the Computed-Tomography (CT) images to diagnose tumor and observe its condition during the treatment process. Due to extensive similarity between pulmonary vessels, bronchus and arteries in lung region and the low contrast of Computed-Tomography (CT) images the accuracy of lung tumor diagnosis is highly dependent on image's contrast and the precision of segmentation. Contrast enhancement and image segmentation are the most prominent image preprocessing techniques that are utilized as a primary and essential steps of almost every pathological applications. Thus, a particular contrast enhancement and image thresholding techniques are required to enhance the contrast of lung CT image by refining their pixels' intensity value and overcome the difficulties of precise segmentation as well as facilitating the accurate pulmonary nodule extraction.

Accordingly, in this research Regional Contrast Enhancement (RCE) and Four-Directional Thresholding (FDT) techniques are introduced followed by nodule extraction and their discrimination based on their respective size and circularity measurements.

Regional Contrast Enhancement (RCE) technique aims to improve the CT image's visual quality by boosting the contrast of lung CT images and modifying the image histogram by implementing the proposed algorithm on every individual pixel based on their intensity value and their regional variations.

The proposed FDT technique also aims to augment the precision of lung CT image's segmentation by implementing a specific thresholding approach from four different directions in which the determination of pixels' value as being either on foreground or background is highly dependent on its adjacent pixel's intensity value and the final decision is made based on all four directions' thresholding results. Finally, pulmonary nodules are extracted from thresholded CT images by several morphological techniques and then extracted candidates are discriminated based on their eccentricity and corresponding size as benign and malignant nodules.

To demonstrate the superiority of proposed RCE technique the minimum Absolute Mean Brightness Error (AMBE), the highest Peak Signal to Noise Ratio and structural Similarity Measurement Index obtained by RCE technique are compared with the other advanced contrast enhancement by histogram equalization methods. The effectiveness and high exactitude of proposed FDT method also has been evaluated on different CT images by correlation and regional non-uniformity measurement criteria. Ultimately, the performance of nodule extraction and discrimination were evaluated and 93.33% of sensitivity, 93.90% of accuracy and 94.59% of specificity have been obtained.



Abstrak tesis yang dikemukakan kepada Senat Universiti Putra Malaysia Sebagai memenuhi keperluan untuk ijazah Master Sains

PENINGKATAN KONTRAS SEKAWASAN DAN TEKNIK-TEKNIK PENGAMBANGAN EMPAT-ARAH BAGI PENGEKSTRAKAN DAN DISKRIMINASI NODUL PULMONARI

Oleh

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Pengekstrakan nodul Pulmonal automatik dan diagnosis penyakit paru-paru menggunakan Sistem Diagnosis Berbantukan Komputer (CAD) merupakan satu tugas yang mencabar. Secara umumnya, sistem CAD menggunakan imej-imej Tomografi-Berkomputer (CT) untuk mendiagnosis tumor dan memerhatikan keadaannya semasa proses rawatan. Oleh kerana persamaan yang banyak di antara pembuluh pulmonari, bronkus dan arteri di kawasan paru-paru dan kontras yang rendah pada imej-imej Tomografi-Berkomputer (CT), ketepatan diagnosis tumor paru-paru adalah sangat bergantung kepada kontras imej tersebut dan ketepatan segmentasinya. Peningkatan Kontras dan segmentasi imej adalah teknik pra pemprosesan imej paling menonjol yang digunakan sebagai langkah utama dan penting bagi hampir setiap aplikasi patologi. Oleh itu, teknik peningkatan kontras dan teknik pengambangan imej tertentu diperlukan untuk meningkatkan kontras imej CT paru-paru dengan memperhalusi nilai keamatan pikselnya dan mengatasi kesukaran mendapatkan segmentasi yang tepat serta memudahkan pengekstrakan nodul pulmonari yang jitu.

Sehubungan itu, dalam kajian ini, Peningkatan Kontras Sekawasan (RCE) dan teknik Pengambangan Empat-arah (FDT) diperkenalkan dan diikuti dengan pengekstrakan dan diskriminasi nodul berdasarkan saiz dan ukuran bundaran masing-masing.

Teknik Peningkatan Kontras Sekawasan (RCE) bertujuan untuk meningkatkan kualiti visual imej CT dengan meningkatkan kontras imej CT paru-paru dan mengubah-suai histogram imej dengan melaksanakan algoritma yang dicadangkan pada setiap piksel individu berdasarkan nilai keamatan mereka dan variasi kawasannya.

Teknik FDT yang dicadangkan juga bertujuan untuk membantu dalam menyokong ketepatan segmentasi imej CT paru-paru dengan melaksanakan pendekatan pengambangan tertentu dari empat arah yang berbeza di mana penentuan nilai piksel sama ada berada di latar depan atau latar belakang adalah sangat bergantung kepada nilai keamatan piksel bersebelahan ini dan keputusan akhir adalah dibuat berdasarkan semua keputusan pengambangan empat arah tersebut. Akhir sekali, nodul pulmonari diekstrak dari imej CT yang telah diambangkan melalui beberapa teknik morfologi dan

kemudiannya imej-imej yang telah diekstrak itu didiskriminasikan berdasarkan keeksentrikan serta saiz padanannya sebagai nodul-nodul tidak merbahaya atau merbahaya.

Untuk menunjukkan keunggulan teknik RCE yang telah dicadangkan, Ralat Kecerahan purata Mutlak (AMBE) minimum, Isyarat Puncak tertinggi kepada Nisbah Bunyi dan Indeks Pengukuran Persamaan struktur yang diperolehi menerusi teknik RCE telah dibandingkan dengan peningkatan kontras termaju lain menerusi kaedah penyamaan histogram. Keberkesanan dan kejituan tinggi yang dipamerkan oleh kaedah FDT yang dicadangkan juga telah dinilai pada imej-imej CT berbeza melalui kaedah korelasi dan kriteria pengukuran bukan keseragaman serantau. Akhirnya, prestasi pengekstrakan dan diskriminasi nodul telah dinilai dan 93.33% sensitiviti, 93.90% kejituan dan 94.59% kespesifikan telah diperolehi.



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I certify that a Thesis Examination Committee has met on 1 July 2015 to conduct the final examination of Saleheh Heidari on her thesis entitled "Regional Contrast Enhancement and Four Directional Thresholding Techniques For Pulmonary Nodule Extraction and Discrimination" in accordance with the Universities and University Colleges Act 1971 and the Constitution of the Universiti Putra Malaysia [P.U.(A) 106] 15 March 1998. The Committee recommends that the student be awarded the Master of Science.

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LIST OF ABBREVIATIONS

ACO	Ant Colony Optimization
AMBE	Absolute Mean Brightness Error
BBHE	Brightness Bi-Histogram Equalization
BW	Black and White image, binary image
CAD	Computer Aided Diagnosis
CAT	Computerized Axial Tomography
CDF	Cumulative Density Function
CLAHE	Contrast Limited Adaptive Histogram Equalization
CT	Computed Tomography
DBS	Discontinuity-Based Segmentation
DICOM	Digital Imaging and Communication in Medicine
DSIHE	Dualistic Sub-Image Histogram Equalization
FDT	Four-Directional Thresholding
FN	False Negative
FP	False Positive
GUI	Graphic User Interface
HE	Histogram Equalization
LCE	Linear Contrast Enhancement
LHE	Local Histogram Equalization
NSCLC	Non-Small Cell Lung Cancer
PDF	Probability Distribution Function
PIV	Pixel's Intensity Value
PNE	Pulmonary Nodule Extraction
PSNR	Peak Signal to Noise Ratio
PSO	Particle Swarm Optimization
RCE	Regional Contrast Enhancement
RMSHE	Recursive Mean Separate Histogram Equalization
RNU	Region Non-Uniformity
ROC	Receiver Operating Characteristic
ROI	Region Of Interest
SBS	Similarity-Based Segmentation
SCLC	Small Cell Lung Cancer
SPN	Solitary Pulmonary Nodule
SSIM	Structural Similarity Index Measurement
T	Thresholding Value
TN	True Negative
ТР	True Positive

C

CHAPTER 1

INTRODUCTION

1.1 Background

Lung is the most vital and complex bilateral organ in human body. It is enclosed by musculoskeletal chest wall and carries out the task of intrathoracic air-exchange by expanding and contracting up to 20 times per minute (Apar K. et al., 2013). Lung is divided into lobes and provides capillaries by supplying oxygen to be diffused to tissue and oxygenate blood all over the body. The healthiness of this respiratory organ is vitally important for human life. Thus it should be retained from any respiratory disorders (American Thoracic Society, 2014).

Recently, many industrialized countries suffer from air pollution that can be the common cause of respiratory diseases such as Acute bronchitic, Lung cancer, Pneumonia, Asthma and Tuberculosis (G. Krucik, 2013). Among all types of lung diseases, Interstitial Lung Diseases (ILD) and Diffuse Parenchymal Lung Disease (DPLD) are generally recognized as the most common and prevalent lung disorders (Guo J. et al., 2002; Helen et al., 2011).

Generally, lung cancer is a type of pulmonary disease that occurs when the anomalous cells grow and proliferated in lung lobes uncontrollably. The nominated abnormal cells usually are divided rapidly and forming the lung pulmonary tumors which diminish the lung's ability to oxygenate the vessels (Peter C., 2009).

Regarding the lung cancer related mortality; National Cancer Institute revealed that there were 226160 new lung cancers diagnosed and 160340 related death by the end of 2012. Accordingly, World Health Organization (WHO) broadcast that the 7.6 million deaths are caused by cancer and by far the lung cancer with 1,370,000 deaths per year is the worst cancer killer worldwide (Peter C., 2009). Studies done in United State manifest that the lifetime risk of invasive lung cancer for women is approximately about 5.5 percent and 5.9 percent for men. It is argued that approximately about 80 percent of lung cancer could be treated if it is diagnosed in incipient phase of disease (Peter J. et al., 2013). Therefore, due to the importance of early disease-diagnosis, the identification of pathological feature and types of pulmonary tumors in lung cancer could impact on accurate prognosis (Russell C. et al., 2008; Rina D. et al., 2013).

Tumors' location variation and their characteristics may differ in specific period of time. Based upon these characteristics and the tumors' traits they are classified into two different types, which determine whether they are cancerous or non-cancerous. The tumors that have not been propagated in specific time and their characteristics remain

identical are known as benign tumors (non-cancerous). Otherwise, they are categorized as malignant tumors as they behave like cancerous nodules that spread to other parts of body through lymphatic system (Peter C., 2009; Robert M. et al., 2014).

It is noticeable that the treatment of lung malignant nodules is much harder and in some case it is impossible. Thus, recognizing pulmonary nodules in early phase of disease has significant impacts on all stages of treatment process (Chaudhary, 2012). Inspection of lung pulmonary disease and identification of tumors' type could be done by scrutinizing Computed-Tomography (CT) images and morphological analysis of extracted nodules.

Lung Computed Tomography (CT) image is a standard instrument, which is utilized for lung disease diagnosis. But due to the specific gray-level distribution among CT images, they have a very low contrast. Although overwhelming minor lung diseases could be observable in CT images, the paucity of information that they provide is often insufficient to identify the type of nodules and hampers the task of pulmonary nodule extraction in Computer Aided Diagnosis (CAD) systems (Guo J. 2002). Lung nodules extraction and malignant tumor detection through Lung CT images investigation by CAD systems has always been a challenging task which is associated with some difficulties. Tackling with these difficulties would not be possible without using imageprocessing techniques.

1.2 Problem Statement

Although body possess natural defenses to protect lung from any germs and large particles such as dust and pollen, air pollutants usually harm the lung tissues directly and undermine those important defenses. Thus, due to the complexity and substantial functioning of lung, the healthiness of this respiratory organ is vitally important for human life and helps the human body to work properly.

Retention of lung from any aforementioned factors is often inevitable. But restraining the development of lung disease like pulmonary malignant and benign nodules and repelling their progression could always prevent lung disease mortality. Consequently, lung Computed-Tomography (CT) image is used to observe lung pulmonary nodules' abnormality (William E. Brant, 2012). Pulmonary nodules are usually emerged as the primary sign of lung cancer, which could be observable in CT images. CT image is the most fundamental tool used in Computer Aided Diagnosis (CAD) system to evaluate the various types of lung lesion (Li Y. et al., 2011). But usually due to narrow dynamic range (poor quality) of CT images, inspection of lung disease and examination of nodules' types become awkward (Guodong Z. et al., 2008; Agarwal, T.K. et al., 2014).

Thus, the low quality of lung CT images is one of the most important obstacles in pulmonary nodule investigation thorough CT images. The poor quality provided by CT image is due to the dynamic range of gray level, the distribution of gray pixels and

their frequency of occurrence throughout the image. This characteristic always causes the lack of contrast in CT image and leads the image to provide paucity of information that makes the nodule examination and their interpretation tedious and difficult.

Additionally, low contrast of lung CT images usually leads to omission of pulmonary nodule recognition by human radiologist. Beside the fact that low contrast of lung CT image make the lung disease diagnosis abstruse for human, it also leads to a high False-Negative (FN) rate for small nodules detection through CAD system (Dolejši, 2007). Observation of pulmonary nodules' abnormalities in CAD system by using lung low contrast CT images is often unfathomable and it will also affect all further processing and analysis. Thus, a contrast enhancement technique is indispensible preprocessing technique to tackle with the lack of contrast in lung CT images.

Extensive similarity between pulmonary vessels, bronchus and arteries in lung region is another factor, which affects pulmonary nodule extraction in CAD system. In spite of the nominated factors, location diversity of lung pulmonary nodules also makes their investigation a difficult task. Therefore, precise image segmentation could be exploited as another preprocessing technique to cope with difficulties involved in pulmonary nodule detection. Although, Image segmentation technique could simply help to extract pulmonary nodule candidates, yet it is a challenging task (Hui Cui, et al., 2013).

Many researchers merely attempted to develop manifold image contrast enhancement and segmentation techniques for general kind of images. Even though these techniques yield adequate results on general images, their performance on lung CT images yields insufficient results. Unexpected results obtained by advanced techniques are due to lung CT images specific characteristics. Thus, in order to cope with lung CT image characteristics a particular image preprocessing is required to tackle with CT images characteristic as well as facilitating and boosting the accuracy of pulmonary nodule extraction.

At last, it is noticeable that the implementation of image processing techniques on lung CT images produces many tumor-like candidates, which could be enumerated as a tumor. So a specific rule is required to identify the type of tumor. In this case, the morphological characteristics of the tumor should be investigated. The features and the traits of the candidates should be analyzed in order to remove the isolate pixels (noise) and eliminate the small candidates that are not identified as a malignant tumor. Therefore, nodules' discrimination and malignant tumor identification is highly challenging.

Ultimately, all aforementioned obstacles that pulmonary nodule detection encounters throughout its performance could simply be summarized as below:

• Dynamic range of gray level, the distribution of gray pixels and their frequency of occurrence throughout the Lung CT images that yield insufficient contrast in CT image (Agarwal, T.K. et al., 2014) and cause

to provide paucity of information that makes the nodule examination and their interpretation difficult.

- Location diversity of lung pulmonary nodules.
- Extensive similarity between pulmonary nodules, vessels, bronchus and arteries in lung region (Hui Cui, et al., 2013).
- Deficiency and unavailability of particular image preprocessing techniques for lung CT images.

1.3 Objectives of Research

As discussed earlier, exactitude and high precision of pulmonary nodule extraction is dependent on several factors. This research is conducted to overcome with these factors by obtaining specific aims as follow:

- To enhance the contrast of lung CT image by proposing a particular image contrast enhancement technique.
- To propose a precise image segmentation (Thresholding) technique to facilitate the pulmonary nodules' candidates extraction.
- To improve the detection of pulmonary nodules without being affected by their location diversity
- To discriminate the extracted candidates as malignant or benign based on their size and eccentricity

1.4 Research Questions

Having an organized mind has always been useful to construct the research substratum comprehensibly that leads the research into a well-projected path. Thus, in order to conduct the research in a well resasoned and coherent way several questions have been designed based on the aformentioned research problems and objectives. The questions are as below:

- Which image processing techniques can be utilized to optimize the lung CT image contrast?
- Which image thresholding technique could be exploited to enhance the precision of lung CT image segmentation?
- How could juxtapleural nodules be extracted as well as parenchymal nodules?
- What type of morphological analysis could sufficiently discriminate the benign and malignant nodules?

1.5 Scope of Research

The scope of this study is to extract and discriminate lung pulmonary nodules solely by image processing technique. The cardinal image processing techniques used in this research are simply divided into image preprocessing such as contrast enhancement and image segmentation and the post processing techniques to extract the nodule candidate and isolate them based on their size and eccentricity as malignant or benign nodules. The instrument used in this research is the original lung Computed-Tomography (CT) image known as lung DICOM files.

The results obtained in image contrast enhancement is compare with six advanced technique such as HE, CLAHE, DSIHE, BBHE, LHE and RMSHE. The performance of image segmentation (Thresholding) technique also has been compared with OTSU and PSO, which are the well-known image segmentation techniques. Furthermore, the performance of pulmonary nodule classification is evaluated by specific criteria to measure it sensitivity and accuracy.

1.6 Contribution of Research

The accuracy of lung nodule extraction and their classification is extremely dependent on the quality of CT images. As stated previously, the quality of lung CT is inappropriate for further analysis. Thus a meticulous image preprocessing is required to simplify the nodule extraction task and enhance it exactitude. Therefore, the major contributions of this research is to propose a method to enhance the contrast of CT images as well as propounding precise image segmentation and nodule extraction.

In this research a Regional Contrast Enhancement (RCE) technique is proposed to boost up the quality and contrast of lung CT images and refine their pixels' intensity value for further analysis. The RCE technique also redistributes the gray pixels and equalized the gray-level's interval to regenerate image histogram. The RCE algorithm is mainly work based on pixels' regional variation throughout the image.

Another contribution of this research is the proposed Four-Directional Thresholding (FDT) techniques in which the neighbor pixels play significant role in determination of every individual pixel's intensity value. Ultimately, nodules will be extract after implementation of those two proposed techniques by performing various morphological techniques. Then extracted nodule will be investigated and discriminated into benign and malignant nodules.

1.7 Thesis Organization

This chapter provides the introduction to the research discussed in this study. The current chapter basically explicates preliminary information about lung nodule extraction. It also provides related problem that had been the main focus of this study that encouraged researching the solitary pulmonary nodule extraction and their classification. This chapter also clarifies research objectives, questions, contribution and the scopes.

Chapter 2 profoundly expounds rudimentary knowledge about lung structure and lung cancer. The chapter also describes nodules and their characteristics. The features of benign and malignant nodule are also elucidated in Chapter 2 as well as their similarities and differences. Computed tomography and DICOM images are also presented.

Following the rudimental information of lung structure and nodules characteristics the prior studies and their achievements are looked over in Chapter 2. These studies are fundamentally construed into two distinct sections as preprocessing and post processing. In which each section thoroughly reviewed pertinent literatures.

Chapter 3 utterly explicates the methodology utilized to conduct this study. It clearly expounds the process followed in this research as well as exploited methodology. The research design, data collection, sampling and instrumentation are simply described. And the designed framework and performance measurements approaches are thoroughly presented.

The proposed approaches to conduct this research are profoundly delineated in Chapter 4. The proposed algorithms for image preprocessing and post processing are discussed in detail and their info-graphic are provided.

Chapter 5 discusses the experimental results of every single proposed algorithm respectively. It clearly elucidated the performance of proposed techniques and evaluates their result based on specific evaluation metrics given in Chapter 4 and the discussed the achievements.

Ultimately the study is concluded in Chapter 6 and the future work is conferred. It clearly draws conclusion of research and describes the possible future works that could be accomplish to enhance this study.

BIBLIOGRAPHY

- Abdullah-Al-Wadud M. A modified histogram equalization for contrast enhancement preserving the small parts in images. IJCSNS Int J Comput Sci, Network Security 2012;12.
- Aedla, Raju; Dwarakish, G.S; Reddy, D.Venkat, "Satellite image contrast enhancement algorithm based on plateau histogram equalization," Region 10 Symposium, 2014 IEEE, vol., no., pp.213,218, 14-16 April 2014, doi: 10.1109/TENCONSpring.2014.6863028
- A.G. Shanbhag, Utilization of Information Measure as a Means of Image Thresholding, CVGIP: Graphical Models and Image Processing, Volume 56, Issue 5, September 1994, Pages 414-419, ISSN 1049-9652, http://dx.doi.org/10.1006/cgip.1994.1037.
- Agarwal, T.K.; Tiwari, M.; Lamba, S.S., "Modified Histogram based contrast enhancement using Homomorphic Filtering for medical images," Advance Computing Conference (IACC), 2014 IEEE International, vol., no., pp.964,968, 21-22 Feb. 2014, doi: 10.1109/IAdCC.2014.6779453
- Al-amri, Salem Saleh, N. V. Kalyankar, and S. D. Khamitkar, "Linear and non-linear contrast enhancement image." Int. J. Comput. Sci. Network Security, vol.10 No.2, February 2010:139-143.
- Alicia M Prater, 2008, Structure and Function of the Lungs, Anatomy And Physiology, http://www.sciences360.com/index.php/structure-and-function-of-the-lungs-17874/ (accessed 2 July 2014).
- American Cancer Society, 2014, Start and spread of lung cancer, Lung Cancer Prevention and Early Detection, http://www.cancer.org/cancer/lungcancernon-smallcell/moreinformation/lungcancerpreventionandearlydetection/lungcancer-prevention-and-early-detection-what-is-lung-cancer (accessed 18 June 2014).
- American Thoracic Society, 2014, ANATOMY AND FUNCTION OF THE NORMAL LUNG, http://www.thoracic.org/clinical/copd-guidelines/forpatients/anatomy-and-function-of-the-normal-lung.php (accessed 29 June 2014).
- Ann Leung and Robin Smithuis, Solitary pulmonary nodule: benign versus malignant, Department of Radiology, Stanford University Medical Center, Stanford, California and the Department of Radiology, Rijnland Hospital, Leiderdorp, the Netherlands, Retrieved 17 August 2013 from http://www.radiologyassistant.nl/en/p460f9fcd50637/solitary-pulmonarynodule-benign-versus-malignant.html
- Apar K. Ganti, David E. Gerber., 2013. Lung Cancer. 1st Ed. Oxford University Press, USA. ISBN: 0199935939, PP: 39.

- Burns, N. and Grove, S. 2001. The practice of nursing research: conduct, critique and utilization (4th ed). W.B.Saunders: Philadelphia, Pennsylvania, USA.
- Cancer Research UK, 2014, Lung cancer risks and causes, http://www.cancerresearchuk.org/about-cancer/type/lung-cancer/about/lungcancer-risks-and-causes (accessed 2 June 2014).
- Chaudhary, A.; Singh, S.S., "Lung Cancer Detection on CT Images by Using Image Processing," Computing Sciences (ICCS), 2012 International Conference on, vol., no., pp.142,146, 14-15 Sept. 2012, doi: 10.1109/ICCS.2012.43.
- Chen Hee Ooi; Kong, N.S.P.; Ibrahim, H., "Bi-histogram equalization with a plateau limit for digital image enhancement," Consumer Electronics, IEEE Transactions on, vol.55, no.4, pp.2072,2080, November 2009, doi: 10.1109/TCE.2009.5373771.
- Chen Hee Ooi; Isa, N.AM., "Adaptive contrast enhancement methods with brightness preserving," Consumer Electronics, IEEE Transactions on, vol.56, no.4, pp.2543,2551, November 2010, doi: 10.1109/TCE.2010.5681139.
- Chen, S., Ramli, A.R.: Preserving brightness in histogram equalization based contrast enhancement techniques. Digit. Signal Process. 14, 413–428 (2004)
- Devin Kowalczyk, 2014, Defining Primary And Secondary, Primary & Secondary Research: Definition, Differences & Methods. Retrieved 3 April 2014 from http://education-portal.com/academy/lesson/primary-secondary-researchdefinition-differences-methods.html#lesson
- Devin Kowalczyk, 2014, Descriptive Research Design: Definition, Examples & Types, http://education-portal.com/academy/lesson/descriptive-research-designdefinition-examples-types.html#lesson (accessed 5 April 2014).
- Diao Liming; Zhang Caiming; Gao Shanshan, "A new algorithm of automatic lung parenchyma segmentation based on CT images," Consumer Electronics, Communications and Networks (CECNet), 2011 International Conference on, vol., no., pp.976, 979, 16-18 April 2011 doi:10.1109/CECNET.2011.5768258
- Dolejši, M. (2007). Detection of Pulmonary Nodules from CT Scans. Center for Machine Perception, Department of Cybernetics Faculty of Electrical Engineering, Czech Technical University. Retrieved 7 June 2014 from http://cmp.felk.cvut.cz/~dolejm1/noduledetection/noduledetection.pdf
- Ebrahimdoost, Y.; Dehmeshki, J.; Ellis, T.S.; Firoozbakht, M.; Youannic, A.; Qanadli,
 S.D., "Medical Image Segmentation Using Active Contours and a Level Set
 Model: Application to Pulmonary Embolism (PE) Segmentation," Digital
 Society, 2010. ICDS '10. Fourth International Conference on, vol., no.,
 pp.269,273, 10-16 Feb. 2010. doi: 10.1109/ICDS.2010.64
- Field, A. (2005). Discovering Statistics Using SPSS (Introducing Statistical Methods S.) (2nd ed.), Sage Publications

- G. Krucik, 2013, Lung, Healthline, http://www.healthline.com/human-bodymaps/lung, (accessed 23 July 2014).
- George Krucik, 2013, Lung, http://www.healthline.com/human-body-maps/lung (accessed 13 June 2014).
- George Schiffman, Jerry R. Balentine, 2013, Solitary Pulmonary Nodule Overview, eMedicineHealth,http://www.emedicinehealth.com/solitary_pulmonary_nodul e/article_em.htm. (accessed 8 June 2014).
- Golchin, M., F. Khalid, L.N. Abdullah and S.H. Davarpanah, 2013. Shadow detection using color and edge information. J. Comput. Sci., 9: 1575-1588.
- Guo, J.; Reinhardt, J.M.; Kitaoka, H.; Zhang, L.; Sonka, M.; McLennan, G.; Hoffman, E.A., "Integrated system for CT-based assessment of parenchymal lung disease," Biomedical Imaging, 2002. Proceedings. 2002 IEEE International Symposium on, vol., no., pp.871,874, 2002. doi: 10.1109/ISBI.2002.1029398
- Guodong Zhang; Peiyu Yan; Hong Zhao; Xin Zhang, "A Contrast Enhancement Algorithm for Low-Dose CT Images Based on Local Histogram Equalization," Bioinformatics and Biomedical Engineering, 2008. ICBBE 2008. The 2nd International Conference on , vol., no., pp.2462,2465, 16-18 May, 2008, doi: 10.1109/ICBBE.2008.948
- HasanulKabir, Abdullah Al-Wadud, and OksamChae, "Brightness Preserving Image Contrast Enhancement Using Weighted Mixture of Global and Local Transformation Functions", International Arab Journal of Information Technology, Vol. 7, No. 4, pp.403-410, 2010.
- HanzeResearch, 2014, Definition and concepts, What is research, University of Applied Sciences. Retrieved 6 August 2014 from https://www.hanze.nl/en/research/researchportal/centre-of-applied-research-and-innovation/art-society/lifelong-learning-in-music/knowledge-base/online-research-coach/pages/what-is-research.aspx?wbc_purpose=B
- Helen, R.; Kamaraj, N.; Selvi, K.; Raja Raman, V., "Segmentation of pulmonary parenchyma in CT lung images based on 2D Otsu optimized by PSO," Emerging Trends in Electrical and Computer Technology (ICETECT), 2011 International Conference on, vol., no., pp.536, 541, 23-24 March 2011 doi: 10.1109/ICETECT.2011.
- Herlihy, B. (2013), Respiratory System, The Human Body in Health and Illness (5 ed.). Saunders. Page 399
- Hui Cui; Xiuying Wang; Fulham, M.; Feng, D.D., "Prior knowledge enhanced random walk for lung tumor segmentation from low-contrast CT images," Engineering in Medicine and Biology Society (EMBC), 2013 35th Annual International Conference of the IEEE, vol., no., pp.6071,6074, 3-7 July 2013, doi: 10.1109/EMBC.2013.6610937

- Jackson, S.L. (2009). Research Methods and Statistics: A Critical Thinking Approach 3rd edition. Belmont, CA: Wadsworth.
- Jeremy P. T. Ward, J. W. (2010). The Respiratory System at a Glance (3 ed.). Wiley-Blackwell.
- Jing Rui Tang, Nor Ashidi Mat Isa, Adaptive Image Enhancement based on Bi-Histogram Equalization with a clipping limit, Computers & Electrical Engineering, Available online 23 June 2014, ISSN 0045-7906, http://dx.doi.org/10.1016/j.compeleceng.2014.05.017.
- Jobin Christ, M.J.a , Parvathi, R.M.S.b Segmentation of medical image using clustering and watershed algorithms (2011) American Journal of Applied Sciences, 8 (12), pp. 1349-1352.
- Jue Lu; Rongqiang Hu, "A new image segmentation method based on Otsu method and ant colony algorithm," Computer Science and Information Processing (CSIP), 2012 International Conference on, vol., no., pp.767, 769, 24-26 Aug. 2012, doi: 10.1109/CSIP.2012.06308966
- Kothar, C. R. (2004). Types of research, Research methodology: An Introduction, Research Methodology: Methods and Techniques (2nd ed.). New Age International.
- Kotkar, Vijay A., and Sanjay S. Gharde, Review of Various Image Contrast Enhancement Techniques, International Journal of Innovative Research in Science, Engineering and Technology, Vol. 2, Issue 7, July 2013, ISSN: 2319-8753.
- K.S. Sim, C.P. Tso, Y.Y. Tan, Recursive sub-image histogram equalization applied to gray scale images, Pattern Recognition Letters, Volume 28, Issue 10, 15 July 2007, Pages 1209-1221, ISSN 0167-8655, http://dx.doi.org/10.1016/j.patrec.2007.02.003.
- Kuada, J. (2012). Research Strategy and Resources, Objectives of your study, Research Methodology: A Project Guide for University Students. Samfundslitteratur.
- Kumar, R. (2012), The research process: Characteristics and requirements, Research Methodology: A Step-by-Step Guide for Beginners (Third edition ed.). SAGE Publications Ltd.
- K Zuiderveld, "Contrast Limited Adaptive Histogram Equalization," Chapter VIII, Graphics Gems IV, P.S. Heckbert (Eds.), Cambridge, MA, Academic Press, (1994), pp. 474-485
- Lee, J. K., Sagel, S. S., Stanley, R. J., & Heiken, J. P. (2006). Computed Body Tomography with MRI correlation (4th Edition ed.). Lippincott Williams & Wilkins. (page 420-554)

- Lim, ShengHoong and Mat Isa, NorAshidi and Ooi, ChenHee and Toh, KennyKalVin, A new histogram equalization method for digital image enhancement and brightness preservation, Signal, Image and Video Processing, 2013, Pages 1-15, ISSN 1863-1703, http://dx.doi.org/10.1007/s11760-013-0500-z.
- Li Yue; Liu Jie; Meng Lingjun, "Suspected pulmonary nodule detection algorithm based on morphology and gray entropy," Computer Science and Automation Engineering (CSAE), 2011 IEEE International Conference on, vol.4, no., pp.103, 108, 10-12 June 2011, doi: 10.1109/CSAE.2011.
- Lung Cancer Health Center, 2014, Understanding Lung Cancer-the Basics, Retrieved 26 may 2014 from http://www.webmd.com/lung-cancer/guide/understanding-lung-cancer-basics
- Lynne Eldridge, 2014, Lung Nodules-Symptomes, Causes, and Diagnosis. http://lungcancer.about.com/od/symptoms/a/Lung-Nodules.htm (accessed 18 June 2014).
- Mathias Prokop, Michael Galanski, Aart J. Van Der Molen, Cornelia Schaefer-Prokop, (2001). Principles of CT, Spiral CT, and Multislice CT. Spiral and Multislice Computed Tomography of the Body (1 ed.). Thieme.
- Melissa Conrad Stöppler, Jay W. Marks, 2014, How common is lung cancer, http://www.medicinenet.com/lung_cancer/page3.htm (accessed 24 June 2014).
- Melissa Conrad Stöppler, Jay W. Marks, 2014, What are the types of lung cancer, http://www.medicinenet.com/lung_cancer/page7.htm (accessed 25 June 2014).
- Mengxing Huang; Wenjiao Yu; Donghai Zhu, "An Improved Image Segmentation Algorithm Based on the Otsu Method," Software Engineering, Artificial Intelligence, Networking and Parallel & Distributed Computing (SNPD), 2012 13th ACIS International Conference on, vol., no., pp.135, 139, 8-10 Aug. 2012. doi: 10.1109/SNPD.2012.26
- Mohsen Keshani, Zohreh Azimifar, Farshad Tajeripour, Reza Boostani, Lung nodule segmentation and recognition using SVM classifier and active contour modeling: A complete intelligent system, Computers in Biology and Medicine, Volume 43, Issue 4, 1 May 2013, Pages 287-300, ISSN 0010-4825
- Most M.M., Craddick, S., Crawford, S., Redican, S., Rhodes, D., Rukenbrod, F., Laws, R. (2003). Dietary quality assurance processes of the DASH-Sodium controlled diet study. Journal of the American Dietetic Association, 103(10): 1339-1346.
- N. Otsu, A Threshold Selection Method from Gray-Level Histograms," Systems, Man and Cybernetics, IEEE Transactions on, vol.9, no.1, pp.62, 66, Jan. 1979. doi: 10.1109/TSMC.1979.4310076

- National Cancer Institute, 2013, Computed Tomography (CT) Scans and Cancer, http://www.cancer.gov/cancertopics/factsheet/detection/CT (accessed 12 July 2014).
- Nema, 2014, A Brief Background of the DICOM Standard, Digital Imaging and Communications in Medicine, Strategic Document & Principal Contacts, Retrieved 2 July 2014 from http://medical.nema.org/dicom/geninfo/Strategy.pdf
- Nick Efford., 2000, Digital Image Processing: A Practical Introduction Using JavaTM. Pearson Education. Retrieved 23 September 2013 from http://www.cs.auckland.ac.nz/courses/compsci773s1c/lectures/ImageProcessi ng-html/topic3.htm.
- Nickfarjam, A. M.; Soltaninejad, S.; Tajeripour, F., "Supervised bi-level thresholding based on Particle Swarm Optimization," Artificial Intelligence and Signal Processing (AISP), 2012 16th CSI International Symposium on, vol., no., pp.370, 373, 2-3 May 2012. doi: 10.1109/AISP.2012.6313775
- Noah Lechtzin, 2014, Control of Breathing, Retrieved 28 June 2014 from http://www.merckmanuals.com/home/lung_and_airway_disorders/biology_of _the_lungs_and_airways/control_of_breathing.html
- Qingping Wang; Hongyu Zhao; Weiwei Wu; Naichang Yuan, "Algorithm for segmentation based on an improved three-dimensional Otsu's thresholding," Computer Science and Network Technology (ICCSNT), 2012 2nd International Conference on, vol., no., pp.1737, 1740, 29-31 Dec. 2012. doi: 10.1109/ICCSNT.2012. 06526256
- Patel, Omprakash, Yogendra PS Maravi, and Sanjeev Sharma. "A Comparative Study of Histogram Equalization Based Image Enhancement Techniques for Brightness Preservation and Contrast Enhancement." arXiv preprint arXiv:1311.4033 (2013).
- Peter Crosta, 2009, How is lung cancer classified, What Is Lung Cancer, Medical News Today, http://www.medicalnewstoday.com/info/lung-cancer/, (accessed 11 July 2014)
- Peter J. Julien MD. CEDARS-SINAI, 2013, CT Lung Screening. Why is Lung Screening Important? http://www.cedars sinai.edu/Patients/Programs-and Services/Imaging-Center/For-Patients/Exams-by-Procedure/CT-Scans/CT-Lung-Screening.aspx (accessed 17 October 2013)
- Phil Rabinowitz, Stephen B. Fawcett, 2014, Collecting and Analyzing Data, Retrieved 3 August 2014 from http://ctb.ku.edu/en/table-of-contents/evaluate/evaluatecommunity-interventions/collect-analyze-data/main
- Qingzhu Wang, Wenwei Kang, Chunming Wu, Bin Wang, Computer-aided detection of lung nodules by SVM based on 3D matrix patterns, Clinical Imaging,

Volume 37, Issue 1, January–February 2013, Pages 62-69, ISSN 0899-7071, http://dx.doi.org/10.1016/j.clinimag.2012.02.003.

- Rafael C. Gonzalez, and Richard E. Woods, "Digital Image Processing", 2nd edition, Prentice Hall, 2002.
- Rao, D.V.; Reddy, L.P., "Contrast Weighted Perceptual Structural Similarity Index for Image Quality Assessment," India Conference (INDICON), 2009 Annual IEEE, vol., no., pp.1,4, 18-20 Dec. 2009, doi:10.1109/INDCON.2009.5409432
- Rina D. Rudyanto, Gorka Bastarrika, Gabriel de Biurrun, Jackeline Agorreta, Luis M. Montuenga, Carlos Ortiz-de-Solorzano, Arrate Muñoz-Barrutia, Individual nodule tracking in micro-CT images of a longitudinal lung cancer mouse model, Medical Image Analysis, Volume 17, Issue 8, December, 2013, Pages 1095-1105, ISSN 1361-8415, http://dx.doi.org/10.1016/j.media.2013.07.002.
- Robert M. Castellan, Richard E. Manrow, 2010, Lung disease fact sheet, Womens health, https://www.womenshealth.gov/publications/our-publications/fact-sheet/lung-disease.html, (accessed 23 July 2014).
- Russell C. Hardie, Steven K. Rogers, Terry Wilson, Adam Rogers, Performance analysis of a new computer aided detection system for identifying lung nodules on chest radiographs, Medical Image Analysis, Volume 12, Issue 3, June 2008, Pages 240-258, ISSN 1361-8415, http://dx.doi.org/10.1016/j.media.2007.10.004.
- Seattle Cancer Care Alliance, 2014, Pulmonary Nodules,Lung Cancer Early Detection and Prevention Clinic. http://www.seattlecca.org/pulmonary-nodules.cfm, (accessed 7 June 2014).
- Shanmugavadivu, P. and Balasubramanian, K. and Muruganandam, A. Particle swarm optimized bi-histogram equalization for contrast enhancement and brightness preservation of images, The Visual Computer, Volume 30, Issue 4, 2014, Pages 387-399, ISSN 0178-2789, http://dx.doi.org/10.1007/s00371-013-0863-8.
- Shengdong Nie; Lihong Li; Yuanjun Wang; Chaofan He; Feng Ji; Jianmei Liang, "A segmentation method for sub-solid pulmonary nodules based on fuzzy c-means clustering,"Biomedical Engineering and Informatics (BMEI), 2012 5th International Conference on, vol., no., pp.169, 172, 16-18 Oct. 2012. doi: 10.1109/BMEI. 2012. 06513127
- Shukui Li; Shaomin Nie. "Image Segmentation Method of Heavy Forgings Based on Genetic Algorithm," Image and Signal Processing, 2009, CISP '09. 2nd International Congress on, vol., no., pp.1, 4, 17-19 Oct. 2009. doi: 10.1109/CISP.2009.5303977
- Sivakumar, S.; Chandrasekar, C., "Lungs image segmentation through weighted FCM," Recent Advances in Computing and Software Systems (RACSS), 2012

International Conference on, vol., no., pp.109, 113, 25-27 April 2012. doi: 10.1109/RACSS.2012. 6212707

- Soong-Der Chen; Ramli, AR., "Contrast enhancement using recursive mean-separate histogram equalization for scalable brightness preservation," Consumer Electronics, IEEE Transactions on, vol.49, no.4, pp.1301,1309, Nov. 2003, doi: 10.1109/TCE.2003.1261233.
- Soong-Der Chen; Ramli, AR., "Minimum mean brightness error bi-histogram equalization in contrast enhancement," Consumer Electronics, IEEE Transactions on, vol.49, no.4, pp.1310,1319, Nov. 2003, doi: 10.1109/TCE.2003.1261234
- Stopher, P. (2012). Collecting, Managing, and Assessing Data Using Sample Surveys (1st ed.). Cambridge University Press.
- Wayne Goddard, S. Melville, (2004). Descriptive Research, Types of Research, Research Methodology: An Introduction (2nd ed.). Juta Academic.
- W. Dean Bidgood, Jr., Steven C. Horii, Fred W. Prior, Donald E. Van Syckle, 1997, Understanding and Using DICOM, the Data Interchange Standard for Biomedical Imaging. http://www.ncbi.nlm.nih.gov/pmc/articles/PMC61235/ (accessed 17 May 2014).
- W. Doyle, "Operation Useful for Similarity-invariant Pattern Recognition", Journal Association for Computing Machinery, vol.9 (2), pp. 259-267, 1962. doi:10.1145/321119.321123.
- Wingerd, B. (2013), The Respiratory System, The Human Body: Concepts of Anatomy and Physiology (3rd ed.). LWW. Page 358-367
- Whitney, C.W., Lind, B.K., Wahl, P.W. (1998). Quality assurance and quality control in longitudinal studies. Epidemiologic Reviews, 20(1): 71-80.

Wilkinson GA, Fraser RG. Roentgenography of the chest. Appl Radiol 1975;4:41-53.

- William E. Brant, C. A. (2012). Section III, Pulmonary, Fundamentals of Diagnostic Radiology (Fourth, in four volumes edition ed.). (J. Pine, Ed.) Philadelphia: LIPPINCOTT WILLIAMS & WILKINS.
- Vinay Kumar and HimaniBansal, "Performance Evaluation of Contrast Enhancement Techniques for Digital Images", International Journal of Computer Science and Technology, Vol. 2, No. 1, pp.23-27, 2011.
- Xiaohong Shen; Yulin Zhang; Fangzhen Li, "An Improved Two-Dimensional Entropic Thresholding Method Based on Ant Colony Genetic Algorithm," Intelligent Systems, 2009. GCIS '09. WRI Global Congress on, vol.1, no., pp.163, 167, 19-21 May 2009. doi: 10.1109/GCIS.2009.96

- Yen JC, Chang FJ, Chang S (1995), "A New Criterion for Automatic Multilevel Thresholding", IEEE Trans. On Image Processing 4 (3): 370-378, ISSN 1057-7149, doi:10.1109/83.366472
- Yeong-Taeg Kim, "Contrast enhancement using brightness preserving bi-histogram equalization," Consumer Electronics, IEEE Transactions on, vol.43, no.1, pp.1,8, Feb 1997, doi: 10.1109/30.580378.
- Yu Wang; Qian Chen; Baomin Zhang, "Image enhancement based on equal area dualistic sub-image histogram equalization method," Consumer Electronics, IEEE Transactions on, vol.45, no.1, pp.68,75, Feb 1999, doi: 10.1109/30.754419.
- Zhenhua Zhang; Ningning Zhou, "A novel image segmentation method combined Otsu and improved PSO," Advanced Computational Intelligence (ICACI), 2012 IEEE Fifth International Conference on, vol., no., pp.583, 586, 18-20 Oct. 2012. doi: 10.1109/ICACI.2012.06463232