

Breast Tumour Visualization Using 3-D Quantitative Ultrasound Methods

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ABSTRACT

Breast cancer is one of the most common cancer types accounting for 29% of all cancer cases. Early detection and treatment has a crucial impact on improving the survival of affected patients. Ultrasound (US) is non-ionizing, portable, inexpensive, and real-time imaging modality for screening and quantifying breast cancer. Due to these attractive attributes, the last decade has witnessed many studies on using quantitative ultrasound (QUS) methods in tissue characterization. However, these studies have mainly been limited to 2-D QUS methods using hand-held US (HHUS) scanners. With the availability of automated breast ultrasound (ABUS) technology, this study is the first to develop 3-D QUS methods for the ABUS visualization of breast tumours. Using an ABUS system, unlike the manual 2-D HHUS device, the whole patient's breast was scanned in an automated manner. The acquired frames were subsequently examined and a region of interest (ROI) was selected in each frame where tumour was identified. Standard 2-D QUS methods were used to compute spectral and backscatter coefficient (BSC) parametric maps on the selected ROIs. Next, the computed 2-D parameters were mapped to a Cartesian 3-D space, interpolated, and rendered to provide a transparent color-coded visualization of the entire breast tumour. Such 3-D visualization can potentially be used for further analysis of the breast tumours in terms of their size and extension. Moreover, the 3-D volumetric scans can be used for tissue characterization and the categorization of breast tumours as benign or malignant by quantifying the computed parametric maps over the whole tumour volume.

Keywords: 3-D imaging, automated breast ultrasound (ABUS), breast cancer, quantitative ultrasound, tumour visualization

1. INTRODUCTION

According to the World Health Organization, every year, about 1.4 million women are diagnosed with breast cancer worldwide, which accounts for 25% of all cancer cases¹. It resulted in 1.68 million cases and about half a million deaths in 2012¹. An early cancer detection and treatment plays a vital role in increasing the survival rate of patients with breast cancer. Currently, mammography is the most common imaging modality for breast cancer screening. However, mammography has two main drawbacks in screening breast cancer: first, it exposes to ionizing radiation that may cause health problems in longitudinal screening and second, its screening ability to identify abnormal structures such as cancerous tumours significantly declines in cases with dense breasts². Considering the high percentage of women with dense breasts (e.g., in United States, over 46% of all women³ and over 74% of women in ages between 40 to 49 years⁴ have dense breasts), the latter limitation of mammography imposes a high risk on monitoring breast cancers. The problem is more serious as there is an increased risk of developing cancer in women with dense breasts⁵.

In this context, *quantitative ultrasound* (QUS) methods provide a promising framework that can non-invasively and inexpensively be used for the early detection of breast cancer in real-time mode⁶. QUS techniques have also been applied to a variety of other tissue characterization applications to detect tissue abnormalities in, for example, the eye⁷, the prostate⁸, and the myocardium⁹.

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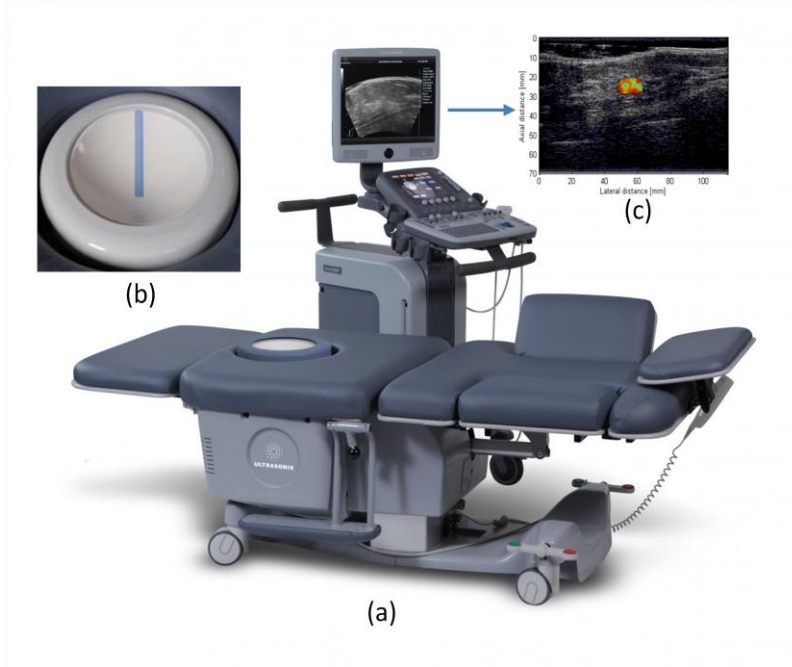


Figure 1- (a) Ultrasonix SonixEmbrace™ automated breast ultrasound (ABUS) System, (b) concave L14-5/115 transducer, (c) two-dimensional parametric map overlaid on the corresponding B-mode image.

Despite recent advances in QUS methods, these techniques are currently limited to 2-D analysis and visualization¹⁰⁻¹⁹. This is mainly because these methods are based on images acquired using hand-held ultrasound (HHUS) devices. Image acquisition using HHUS, however, is relatively time consuming and highly dependent on the experience of the sonographer, in addition to providing only a partial 2-D view of the breast. These limitations have recently been mitigated by the introduction of automated breast ultrasound (ABUS) technologies²⁰⁻²³. ABUS performs automatic, therefore, operator-independent image acquisition from each breast in relatively short time (< 2 minutes). It also enhances the localization of tumour in 3-D within the breast.

In this work, for the first time, ABUS system was used in conjunction with QUS spectral and backscatter coefficient (BSC) methods to provide a 3-D visualization of breast tumours. This can potentially improve the performance of a tissue characterization system based on the QUS methods as a true volumetric analysis of the tumour becomes feasible.

2. METHODS

2.1 Ultrasound data acquisition

Data was acquired from 10 patients over the last 4 months. Ultrasound B-mode and raw radiofrequency (RF) data were acquired using a SonixEmbrace™ (Ultrasonix Med. Corp., Richmond, BC, Canada) ABUS system (Figure 1a) equipped with a concave L14-5/115 transducer (Figure 1b) with the centre frequency of 5.8 MHz and an adjustable imaging bed. The patient lay on the bed in prone position with the breast resting in the imaging dome. The custom concave US transducer embedded into the imaging dome rotated around the entire breast by an automated step motor capturing 154 2-D frames with a known angular radial distance.

2.2 Quantitative ultrasound analysis

The analysis of ultrasound RF data was performed across all frames with identifiable tumour regions by selecting a region of interest (ROI) containing the tumour. Power spectra were calculated using a Fourier transform of the raw radiofrequency data for each scan line through the ROI using a sliding window method. Data were normalized with the averaged power spectrum obtained from an agar-embedded glass-bead phantom model²⁴. Linear regression analysis was

performed on the averaged power spectrum within a central-frequency based -12 dB window to generate a best-fit line²⁵. Three *spectral* parametric maps were subsequently computed including the normalized power at the centre frequency termed as mid-band fit (MBF), spectral intercept (SI) as the intercept of the fit line to the vertical axis, and spectral slope (SS), which was the slope of the fit line. The same normalized power spectrum was used to estimate the *backscatter coefficient* (BSC) parameters over the ROIs using the reference phantom technique²⁶. Subsequently, by a least-squares fitting of the Gaussian form factor to the BSC, parameters of this form factor were computed²⁷, which were corresponding to the maximum coefficient of determination. The parameters computed were including effective acoustic concentration (EAC) and effective scatterer diameter (ESD).

2.3 3-D Visualization

The concave transducer of the ABUS system scanned the breast by rotating around the centre of the dome (see Figure 1b), collecting the frames radially. Therefore, the image acquisition was essentially performed in polar coordinates. Hence, for a 3-D rendering of the parametric maps, the frames could not simply stack on top of each other, and a conversion from polar to Cartesian coordinate was required as a pre-processing step. The conversion was performed with the known angular space between the successive frames as well as the known radius of the dome/transducer. Since the frames were acquired in discrete angular positions, an interpolation was needed as an intermediate step to obtain all X, Y, and Z coordinates of the entire tumour volume. The resulting 3-D matrix was eventually visualized using a transparent color-coded rendering code to enable observation of different layers of the entire tumour all at once.

3. RESULTS

The study involved 10 patients from the ages of 31 to 67 ($53 \pm 12/\text{mean} \pm \text{SD}$), among which two patients were confirmed as having malignant masses and the other 8 as benign cases. Figure 2 displays the results of 3-D visualization of a representative breast tumour using the proposed method for the spectral and BSC parametric maps. The size of tumour and its extension can be read from the axes. The colour bar indicates the range of changes for each parametric map. A transparent rendering was performed to enable visualization of the tumour in all layers at once. Tumours appeared less defined in terms of their borders compared to benign cases. Benign cases also demonstrated more homogenous internal structure compared to tumours.

4. CONCLUSIONS

In this study, automated breast cancer (ABUS) ultrasound machine was used, for the first time, in conjunction with quantitative ultrasound methods to provide a 3-D volumetric visualization of breast tumours. The entire tumour was visualized using a transparent color-coded QUS spectral and backscatter coefficient parametric map. Consequently, the changes in QUS parametric maps could be visualized inside the tumour, which could potentially ease the subsequent analysis of the tumour. In future work, we will provide the visualization of the 3-D parametric maps within the B-mode images. Moreover, the volumetric parametric maps will be rendered in three different planes including sagittal, axial, and coronal planes overlaid on the B-mode images of the corresponding frames. A moving bar will facilitate displaying the planes at any desired point inside the tumour. The 3-D volumetric tumour could also be further analyzed for tissue characterization and discrimination between benign and malignant tumour types.

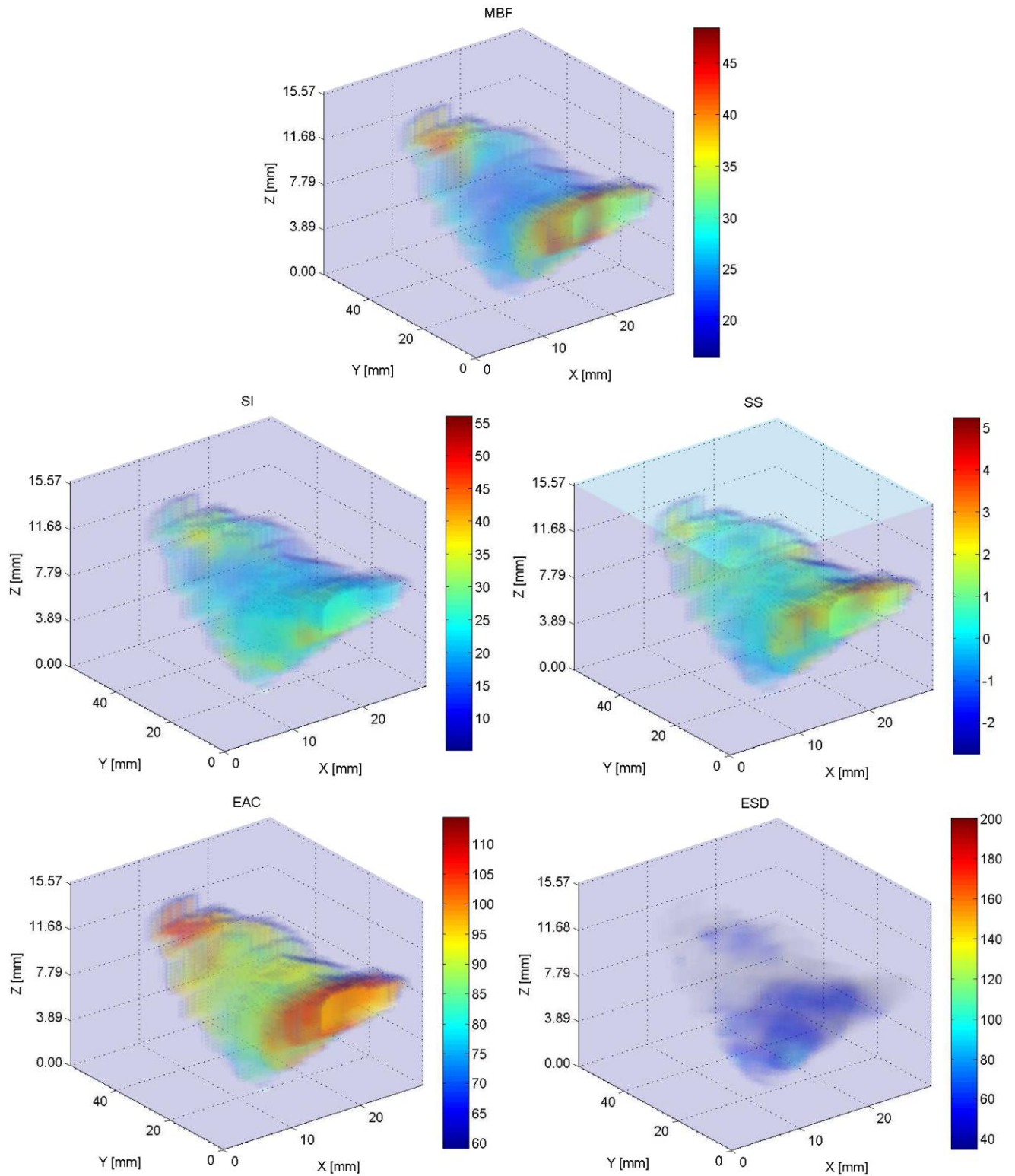


Figure 2-Three-dimensional rendering of a breast tumour using spectral and backscatter coefficient parametric maps computed on the ROIs selected from the frames acquired using an ABUS system.

REFERENCES

- [1] Stewart, B., and Wild, C.P., [World Cancer Report 2014] , International Agency for Research on Cancer (2014).
- [2] Tilanus-Linthorst, M.M., Obdeijn, I.M., Bartels, K.C., de Koning, H.J., and Oudkerk, M., “First experiences screening women at high risk for breast cancer with MR imaging,” *Breast Cancer Research and Treatment* 63(1), 53–60 (2000).
- [3] Pisano, E.D., Constantine, G., Hendrick, E., Yaffe, M., Baum, J.K., Acharyy, S., Conant, E.F., Fajardo, L.L., Bassett, L., et al., “Diagnostic Performance of Digital versus Film Mammography for Breast-Cancer Screening,” *The New England Journal of Medicine* 353(17), 1773–1783 (2005).
- [4] Checka, C.M., Chun, J.E., Schnabel, F.R., Lee, J., and Toth, H., “The Relationship of Mammographic Density and Age: Implications for Breast Cancer Screening,” *American Journal of Roentgenology* 198(3), W292–W295 (2012).
- [5] Boyd, N., Guo, H., and Martin, L., “Mammographic density and the risk and detection of breast cancer,” *The England Journal of Medicine* 356(3), 227–236 (2007).
- [6] Liao, Y.-Y., Tsui, P.-H., Li, C.-H., Chang, K.-J., Kuo, W.-H., Chang, C.-C., and Yeh, C.-K., “Classification of scattering media within benign and malignant breast tumors based on ultrasound texture-feature-based and Nakagami-parameter images,” *Medical Physics* 38(4), 2198 (2011).
- [7] Coleman, D.J., Lizzi, F.L., Silverman, R.H., Helson, L., Torpey, J.H., and Rondeau, M.J., “A model for acoustic characterization of intraocular tumors,” *Investigative Ophthalmology and Visual Science* 26(4), 545–550 (1985).
- [8] Feleppa, E., Kalisz, A., Sokil-Melgar, J., Lizzi, F., Liu, T., Rosado, A., Shao, M., Fair, W., Wang, Y., et al., “Typing of prostate tissue by ultrasonic spectrum analysis,” *IEEE Transactions on Ultrasonics, Ferroelectrics and Frequency Control* 43(4), 609–619 (1996).
- [9] Yang, M., Krueger, T.M., Miller, J.G., and Holland, M.R., “Characterization of Anisotropic Myocardial Backscatter Using Spectral Slope, Intercept and Midband Fit Parameters,” *Ultrasonic Imaging* 29(2), 122–134 (2007).
- [10] Sadeghi-Naini, A., Falou, O., Zubovits, J., Dent, R., Verma, S., Trudeau, M.E., Boileau, J.F., Spayne, J., Iradji, S., et al., “Quantitative Ultrasound Evaluation of Tumour Cell Death Response in Locally Advanced Breast Cancer Patients Receiving Chemotherapy,” *Clinical Cancer Research* 19(8), 2163–2174 (2013).
- [11] Banihashemi, B., Vlad, R., Debeljevic, B., Giles, A., Kolios, M.C., and Czarnota, G.J., “Ultrasound imaging of apoptosis in tumor response: novel preclinical monitoring of photodynamic therapy effects,” *Cancer research* 68(20), 8590–6 (2008).
- [12] Gangeh, M.J., Sadeghi-Naini, A., Kamel, M.S., and Czarnota, G.J., “Assessment of cancer therapy effects using texton-based characterization of quantitative ultrasound parametric images,” in *Proc. Int. Symp. Biomed. Imaging*, 1372–1375 (2013).
- [13] Gangeh, M.J., Tadayyon, H., Sannachi, L., Sadeghi-Naini, A., and Czarnota, G.J., “Quantitative Ultrasound Spectroscopy and a Kernel-Based Metric in Clinical Cancer Response Monitoring,” in *Proc. Int. Symp. Biomed. Imaging From Nano to Macro*, 255–259 (2015).
- [14] Gangeh, M.J., Sadeghi-Naini, A., Diu, M., Tadayyon, H., Kamel, M.S., and Czarnota, G., “Categorizing Extent of Tumour Cell Death Response to Cancer Therapy Using Quantitative Ultrasound Spectroscopy and Maximum Mean Discrepancy,” *IEEE Transactions on Medical Imaging* 33(6), 1390–1400 (2014).
- [15] Gangeh, M.J., Tadayyon, H., Sannachi, L., Sadeghi-Naini, A., Tran, W.T., and Czarnota, G., “Computer Aided Theragnosis Using Quantitative Ultrasound Spectroscopy and Maximum Mean Discrepancy in Locally Advanced Breast Cancer,” *IEEE Transactions on Medical Imaging* In Press, (2016).
- [16] Gangeh, M.J., El Kaffas, A., Hashim, A., Giles, A., and Czarnota, G.J., “Advanced Machine Learning and Textural Methods in Monitoring Cell Death Using Quantitative Ultrasound Spectroscopy,” in *Proc. Int. Symp. Biomed. Imaging From Nano to Macro*, 646–650 (2015).

- [17] Sadeghi-Naini, A., Zhou, S., Gangeh, M.J., Jahedmotlagh, Z., Falou, O., Ranieri, S., Azrif, M., Giles, A., and Czarnota, G.J., "Quantitative evaluation of cell death response in vitro and in vivo using conventional-frequency ultrasound," *Oncoscience* 2(8), 716–726 (2015).
- [18] Gangeh, M.J., Hashim, A., Giles, A., and Czarnota, G.J., "Cancer therapy prognosis using quantitative ultrasound spectroscopy and a kernel-based metric," in *Proc. SPIE Med. Imaging* 9034, 903406(1)–903406(6) (2014).
- [19] Vlad, R.M., Alajez, N.M., Giles, A., Kolios, M.C., and Czarnota, G.J., "Quantitative ultrasound characterization of cancer radiotherapy effects in vitro," *International journal of radiation oncology, biology, physics* 72(4), 1236–43 (2008).
- [20] Giuliano, V., and Giuliano, C., "Using Automated Breast Sonography as Part of a Multimodality Approach to Dense Breast Screening," *Journal of Diagnostic Medical Sonography* 28(4), 159–165 (2012).
- [21] Zhuang, B., Chen, T., Leung, C., Chan, K., Dixon, J., Dickie, K., and Pelissier, L., "Microcalcification Enhancement in Ultrasound Images from a Concave Automatic Breast Ultrasound Scanner," in *Proc. IEEE Int. Ultrason. Symp.*, 1662–1665 (2012).
- [22] Azar, R.Z., Leung, C., Chen, T.K., Dickie, K., Dixon, J., Chan, K.-K., and Pelissier, L., "An automated breast ultrasound system for elastography," in *Proc. IEEE Int. Ultrason. Symp.*, 1–4 (2012).
- [23] OK-Chen, T.K., Leung, C., Azar, R.Z., Chan, K.-K., Zhuang, B., Dickie, K., Dixon, J., Pendziwol, L., and Pelissier, L., "Importance of Transducer Position Tracking for Automated Breast Ultrasound: Initial Assessments," in *Proc. IEEE Int. Ultrason. Symp.*, 2623–2626 (2012).
- [24] Dong, F., Madsen, E.L., MacDonald, M.C., and Zagzebski, J.A., "Nonlinearity parameter for tissue-mimicking materials," *Ultrasound in Medicine & Biology* 25(5), 831–838 (1999).
- [25] Lizzi, F.L., "Ultrasonic spectrum analysis for tissue assays and therapy evaluation," *International Journal of Imaging Systems and Technology* 8(1), 3–10 (1997).
- [26] Yao, L.X., Zagzebski, J.A., and Madsen, E.L., "Backscatter coefficient measurements using a reference phantom to extract depth-dependent instrumentation factors," *Ultrasonic imaging* 12(1), 58–70 (1990).
- [27] Insana, M.F., and Hall, T.J., "Parametric ultrasound imaging from backscatter coefficient measurements: Image formation and interpretation," *Ultrasonic Imaging* 12(4), 245–267 (1990).