

Diffraction and coherence in breast ultrasound tomography: A study with a toroidal array

F. Simonetti^{a)}

Department of Mechanical Engineering, Imperial College, London SW7 2AZ, United Kingdom
and MS D443, Los Alamos National Laboratory, Los Alamos, New Mexico 87545

L. Huang

MS D443, Los Alamos National Laboratory, Los Alamos, New Mexico 87545

N. Duric and P. Littrup

Karmanos Cancer Institute, Wayne State University, 4100 John R, Detroit, Michigan 48201

(Received 17 July 2008; revised 11 May 2009; accepted for publication 13 May 2009;
published 9 June 2009)

Ultrasound is commonly used as an adjunct to mammography for diagnostic evaluation of suspicions arising from breast cancer screening. As an alternative to conventional sonography that uses hand-held transducers, toroidal array probes that encircle the breast immersed in a water bath have been investigated for ultrasound tomography. In this paper, two sets of experiments performed with a prototype ultrasound scanner on a phantom and a human breast *in vivo* are used to investigate the effects of diffraction and coherence in ultrasound tomography. Reconstructions obtained with transmission diffraction tomography (TDT) are compared with conventional reflection imaging and computerized ultrasound tomography showing a substantial improvement. The *in vivo* tests demonstrate that TDT can image the complex boundary of a cancer mass and suggest that it can reveal the anatomy of milk ducts and Cooper's ligaments. © 2009 American Association of Physicists in Medicine. [DOI: 10.1118/1.3148533]

Key words: ultrasound, tomography, breast, imaging

I. INTRODUCTION

Since the 1970s, researchers have been exploring the potential of ultrasound tomography for the detection of breast cancer.¹⁻³ Indeed, ultrasound could be a very attractive alternative to the gold standard of X-ray mammography since it is intrinsically safer, it can detect some cancers that are invisible on mammograms and could lead to earlier diagnosis.⁴ In addition, ultrasound technology is inexpensive compared to x ray or MRI, thus offering the potential for the development of a dense network of screening centers that could deliver diagnostic capabilities to the bulk of the population.

While the ray model of geometrical optics is, in general, sufficient to describe the propagation of high energy photons in X-ray tomography of soft media; diffraction, refraction, and scattering can become dominant when imaging complex objects, such as the human breast, with ultrasound. Although the anatomy of the breast is not yet completely understood,⁵ it is largely accepted that the gross anatomy consists of a fatty tissue and a glandular component which is arranged in lobes that are held together by a fibrous framework called Cooper's ligaments. The lobes contain lobules that are connected to the nipple by an entangled network of ducts which Cooper, in the celebrated *On the Anatomy of the Breast*,⁶ compared to the intertwined root of a tree. These structures along with lymphatic and blood vessels result in the very complex anatomy of the breast. When an ultrasonic wave travels through breast tissue, the mechanical properties of these structures produce a rich spectrum of phenomena that change the characteristics of the ultrasound field. The extent

of these changes depends on the characteristic size of these structures relative to the wavelength λ of the probing wave. As an example, randomly distributed features on the scale of λ can result in complex interference patterns that cause the granular appearance of conventional sonograms known as speckle and which do not bear any obvious relationship to the actual anatomy.⁷ Similarly, multiply scattered sound between strong, closely spaced, and randomly distributed features can cause the ballistic wave transmitted through a complex medium to be followed by a complex coda statistically described by diffusion equations.^{8,9}

Recent progress in solid state electronics and increased computational power have reinvigorated interest in ultrasound tomography of the breast, and much effort is now being devoted to the development of a new generation of ultrasound scanners that employ a toroidal array of sensors¹⁰⁻¹³ as depicted in the diagram of Fig. 1(a). While in the early work by Carson *et al.*,³ the breast was probed by means of a pair of transducers mechanically scanned around a circular aperture that encircled the breast, the toroidal array replaces the mechanical scanning with a much faster electronic switching that allows the breast to be insonified from any direction in the plane of the aperture and the scattered field to be measured all around the breast in less than 0.1 s. Note that the speed at which the measurements are performed is crucial to avoid artifacts caused by tissue motion.

This full view configuration can be used to perform transmission measurements through the breast in conjunction with the more conventional backscattering measurements ob-

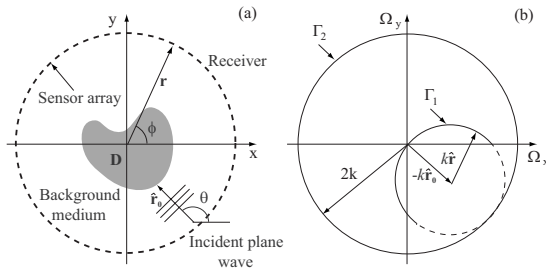


FIG. 1. (a) Diagram of a toroidal array used in ultrasound tomography. (b) Two dimensional K -space showing how the scattered field measured in the direction $\hat{\mathbf{r}}$ and due to an incident plane wave from direction $\hat{\mathbf{r}}_0$ maps onto the point $\Omega = 2\pi/\lambda(\hat{\mathbf{r}}_0 - \hat{\mathbf{r}})$ of the K -space.

tained with linear array probes. The combination of backscattering and transmission measurements can then be used to extract accurate information about the speed of sound and attenuation distributions within the breast. Backscattering measurements are used to reproduce compound beam-forming³ synthetically and lead to the so-called reflection image (RI). A subset of the array sensors is used to create a virtual aperture that produces different images of the breast, which are then combined together as the aperture revolves around the breast.¹³ On the other hand, transmission measurements are used in computerized ultrasound tomography (CUT) to estimate the arrival time of the ballistic transmission through the breast, that is, subsequently used to reconstruct the speed of sound map via the Radon transform.¹⁴ Promising results for human breast *in vivo* have been reported by Carson *et al.*³ and more recently by Duric *et al.*¹³ with a toroidal array. However, CUT is based on the ray approximation that is known to cause resolution degradation and image artifacts due to its inability to account for diffraction.¹⁵ Therefore, it can be expected that more accurate reconstruction methods such as diffraction tomography¹⁶ (DT) could lead to improved resolution. This effect has already been observed by Devaney¹⁷ when comparing the backprojection method that is based on the ray theory and the backpropagation¹⁸ method that accounts for diffraction. An improvement on image resolution is desirable since it would decrease the size of the minimum detectable lesion and increase the specificity of CUT by providing additional anatomical information.

The aim of this paper is to investigate the role of diffraction and coherence in ultrasound breast tomography by means of two sets of experiments performed on a phantom and a human breast *in vivo*. The study on the effects of diffraction employs the DT approach introduced in Ref. 19 and has a twofold objective. First, it is to assess if transmission diffraction tomography (TDT), which uses the far receiver semiarc of the array, leads to higher image resolution than CUT. Second, it is to investigate, in two examples, whether RI, which uses the near receiver semiarc of the array, and TDT show evidence of offering complementary information. In fact, it is known that RI and CUT images complement each other since the former provides information about the boundaries of regions of sudden impedance variations within

the object, whereas, the latter supplies quantitative information about the spatial impedance variations.³

While CUT requires the use of wideband (WB) insonification to time the journey of ultrasound through the breast, RI and DT can also be performed with continuous wave (cw) excitation. This is because both RI and DT can be seen as interferometric techniques that exploit spatial interference caused by waves transmitted and detected by transducers at different spatial locations. A cw image is coherent and can contain speckle. Although speckle contrast is used in sonograms to detect different structures, speckle can mask small lesions, thus reducing the sensitivity of sonography. Therefore, the question arises whether or not speckle noise reduction is beneficial. Here, in order to address this question speckle reduction is obtained by incoherent superposition of different cw images, within the bandwidth of a WB insonification. Incoherent superposition is known to reduce speckle in the same way as incoherent light cancels speckle noise in optical imaging.²⁰

II. METHODS

We conduct ultrasound tomography experiments on a phantom and a human breast *in vivo* to investigate the effects of diffraction and coherence in breast tomography. While the knowledge of the true *in vivo* anatomy is always uncertain, the properties of a phantom can be characterized accurately during its manufacturing or by means of imaging techniques that cannot be used *in vivo*, such as X-ray CT. The knowledge about the true structure of the phantom can then be used to benchmark different ultrasound imaging methods. However, *in vivo* validation of observations carried out on a phantom is paramount due to the complex biology of the actual breast.

In this paper, imaging methods based on different physical models are applied to the two experimental data sets, and the resulting reconstructions are compared against each other.

II.A. Physical model

The objective of ultrasound tomography is to reconstruct the spatial distribution of a target material property from the perturbation induced by the object's structure on the free propagation of ultrasound. Such a material property is described by the object function that is related to the index of refraction map, $n(\mathbf{r})$, via the relationship $O(\mathbf{r}) = k^2[n(\mathbf{r})^2 - 1]$, where k is the wavenumber of the probing wave in the free space. In this paper energy dissipation phenomena are neglected and it is assumed that n is real. This assumption is valid as we consider relatively low frequency where the effect of material absorption is negligible compared to the attenuation caused by scattering. For DT, central to the reconstruction is the existence of a one-to-one mapping between the perturbation p and the spatial Fourier transform of the object function O , which defines the so-called K -space,

$$p \leftrightarrow O. \tag{1}$$

The definition of the perturbation in Eq. (1) depends on the model used to describe the interaction between the incident wave and the probed object.

To illustrate this, let us consider the two-dimensional scattering problem depicted in Fig. 1(a), whereby a monochromatic (cw) plane wave propagating in the direction $\hat{\mathbf{r}}_0$ is incident on an object of support D . Assuming that the scattering problem can be expressed by a scalar potential, the field detected by an array sensor placed in the far field at position \mathbf{r} , $\psi(\mathbf{r}, k\hat{\mathbf{r}}_0)$, is given by

$$\lim_{r \rightarrow \infty} \psi(\mathbf{r}, k\hat{\mathbf{r}}_0) = \exp(ik\hat{\mathbf{r}}_0 \cdot \mathbf{r}) + f(k\hat{\mathbf{r}}, k\hat{\mathbf{r}}_0) \frac{\exp(ikr)}{\sqrt{r}}, \tag{2}$$

where the first term of the right hand side is the incident plane wave, which propagates with wavelength λ ($k = 2\pi/\lambda$) and $f(k\hat{\mathbf{r}}, k\hat{\mathbf{r}}_0)$ is the scattering amplitude defined as

$$f(k\hat{\mathbf{r}}, k\hat{\mathbf{r}}_0) = \Pi \int_D d^2r' \exp(-ik\hat{\mathbf{r}} \cdot \mathbf{r}') O(\mathbf{r}') \psi(\mathbf{r}', k\hat{\mathbf{r}}_0), \tag{3}$$

with

$$\Pi = \frac{\exp(i\pi/4)}{\sqrt{8\pi k}}. \tag{4}$$

It can be shown that under the Born approximation, the perturbation p in Eq. (1) coincides with the scattering amplitude²¹

$$p(k\hat{\mathbf{r}}, k\hat{\mathbf{r}}_0) = f(k\hat{\mathbf{r}}, k\hat{\mathbf{r}}_0) = \Pi O[k(\hat{\mathbf{r}} - \hat{\mathbf{r}}_0)]. \tag{5}$$

This relationship links the measurements to the Fourier transform of the object function $O(\Omega)$ directly. In fact, the scattering amplitude can be measured experimentally with a toroidal array of transreceivers, by transmitting with each sensor sequentially and detecting the field with all the sensors in parallel.

The diagram in Fig. 1 shows how a particular transmit and receive pair maps onto a point of the K -space. In particular, for a given transmitter position defined by $\hat{\mathbf{r}}_0$ the measurements map onto the circle Γ_1 as the receiver direction $\hat{\mathbf{r}}$ spans the entire array. This is known as the Ewald circle. The solid part of the circle corresponds to the transmission measurements (the angle between $\hat{\mathbf{r}}$ and $\hat{\mathbf{r}}_0$ is less than $\pi/2$), whereas the dashed part corresponds to the back-scattering measurements. As the position of the transmitter revolves around the object, the Ewald circle sweeps a disk of the K -space with radius $2k$ known as the Ewald limiting disk (ELD) and labeled as Γ_2 in Fig. 1(b).

It has to be stressed that the validity of Eq. (5) depends on the assumptions that the Born approximation applies and that the incident field is a plane wave. The Born approximation can be considered to be valid when material contrast is low and the object is small compared to the wavelength.²²⁻²⁶ In particular, it is found that these conditions are met when the object function satisfies the relationship

$$\sup_{r < l} |O(\mathbf{r})| < c \frac{\lambda}{l}, \tag{6}$$

where l is the characteristic size of the object and c is a constant for which different values have been proposed.²⁵⁻²⁷ Equation (6) has an important physical interpretation. Although the incident wavefront becomes distorted as it travels through the object, the phase delay accumulated as the wave passes through the object is always much smaller than π . This leads to a relatively smooth wavefront emerging from the object as destructive and constructive interference phenomena cannot occur. However, this is not true in the case of the human breast at the standard submillimeter wavelengths used in practice. Indeed strong interference phenomena that lead to complex refraction patterns have been observed.^{28,29} On the other hand, the refraction phenomenon can be exploited to isolate information relative to the targeted region inside the breast according to the multiscale method introduced in Ref. 19 and briefly discussed in Sec. II D.

In practice it is not possible to produce a pure plane wave. However, if a small source is placed at a distance from the object much larger than l , the wave front incident on the object can be considered locally planar. If the active surface of the source is an area smaller than $\lambda \times \lambda$ the radiated wave field can be considered as a spherical wave. A quasicylindrical wavefront can be produced with a rectangular active area with one side smaller than λ and the other much larger. This configuration is that used in the experiments reported in this paper and generates a quasicylindrical wave in the x - y plane in Fig. 1(a).

II.B. Imaging methods

From the knowledge of $O(\Omega)$ within the ELD, $O(\mathbf{r})$ can be reconstructed by assuming that $O(\Omega)$ vanishes outside the ELD and performing the inverse Fourier transform. This leads to a low-pass filtered image of the object function, I_{DT} , which in the K -space is given by

$$I_{\text{DT}}(\Omega) = O(\Omega) H_{\text{DT}}(\Omega), \tag{7}$$

with

$$H_{\text{DT}}(\Omega) = \begin{cases} 1 & |\Omega| < 2k \\ 0 & |\Omega| > 2k. \end{cases} \tag{8}$$

This implies that only the characteristics of the object that vary on a spatial scale longer than $\lambda/2$ can be reconstructed, leading to the classical diffraction limit.³⁰

In transmission diffraction tomography (TDT), the object function is reconstructed from only the transmission measurements. Therefore, as the transmitter revolves around the object, the half solid circle in Fig. 1(b) describes a disk of radius $\sqrt{2}k$ centered at the origin and contained within the ELD. In other words, TDT provides a low-pass filtered image with cutoff $\sqrt{2}k$ rather than $2k$. A similar argument shows that reflection diffraction tomography (RDT) provides a band-pass filtered image of the object with cutoffs at $\sqrt{2}k$ and $2k$. This analysis suggests that RDT complements TDT by reconstructing the spatial frequencies between $\sqrt{2}k$ and $2k$.

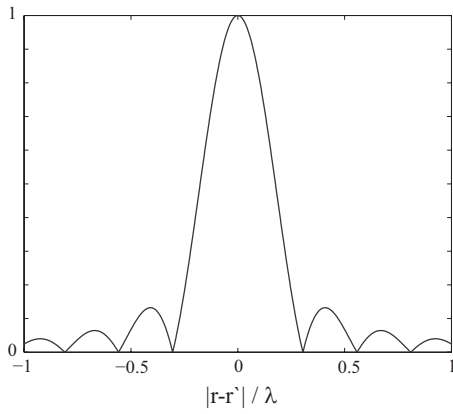


FIG. 2. Normalized modulus of the point spread function for 2D diffraction tomography.

RDT and TDT can be related to reflection imaging (RI) and computerized ultrasound tomography (CUT), respectively. RI can be seen as a beam forming process that provides an image of the object by focusing and steering a beam across the object. As shown in Ref. 31, the beam forming image is a filtered version of the DT reconstruction. The filter introduces a distortion of the DT image by attenuating the spatial frequencies in the order of k and amplifying those approaching $2k$.

CUT can be considered as the limit of TDT obtained when the wavelength vanishes. This asymptotic condition is achieved in practice when the wavelength is much smaller than the spatial scale over which variations of the object function occur, leading to the approximation of geometrical optics. In this case, k approaches to infinity and the half circle in the K -space representing the transmission measurements, Fig. 1(b), tends to a straight line passing through the origin, leading to the Fourier slice theorem.¹⁶

II.C. Frequency diversity

The previous analysis has considered cw excitation only; therefore, it refers to the perfectly coherent case. It can be observed that the imaging process is the convolution of the object function with the point spread function (PSF) h ,

$$I(\mathbf{r}, \omega) = \int_D d^2r' h(\mathbf{r} - \mathbf{r}', \omega) O(\mathbf{r}'), \quad (9)$$

where the PSF is the image of a point as seen by a particular imaging method and it is a function of frequency. As an example, in the case of DT, the PSF is the inverse Fourier transform of Eq. (8) i.e.,

$$h_{DT}(\mathbf{r}, \mathbf{r}') = k \frac{J_1(2k|\mathbf{r} - \mathbf{r}'|)}{\pi|\mathbf{r} - \mathbf{r}'|}, \quad (10)$$

where J_1 is the order one Bessel function of the first kind. Figure 2 shows the modulus of the PSF as a function of the distance between a point in the object \mathbf{r}' and a point in the image \mathbf{r} relative to λ . The presence of a main central lobe means that only the points of the object contained within the main lobe give a significant contribution to the integral in

Eq. (9). Therefore, the image in \mathbf{r} is given by the coherent superposition of $O(\mathbf{r}')$ within the main lobe of the PSF. If $O(\mathbf{r}')$ varies randomly within the main lobe, then the integral in Eq. (9) is a random function of \mathbf{r} . Moreover, at a single position \mathbf{r} , the image does not relate to the object function at a particular point in space. These are the main implications of coherence and explain the speckle phenomenon.

In optics it is known that speckle can be suppressed by reducing the temporal coherence of the illumination²⁰ and thus a WB source rather than a monochromatic one should be used. In fact, optical detectors average the intensity of light over a time interval that is much longer than the characteristic period of light wavepackets. This means that in optics a WB image is given by the incoherent superposition of all the monochromatic images, $I(\mathbf{r}, \omega)$, that could be reconstructed if it was possible to separate the different frequency components of the illumination,

$$I_{\text{incho}}(\mathbf{r}) = \int_{-\infty}^{\infty} d\omega |I(\mathbf{r}, \omega)|^2. \quad (11)$$

Note that the superposition is incoherent because the phase information of each monochromatic image is not included in Eq. (11). On the other hand, ultrasonic sensors generate an almost instantaneous measurement of the wave field, thus allowing the harmonics contained in the insonification to be separated. As a result, RI with WB excitation is still a coherent process. In fact, the wideband version of Eq. (9) becomes

$$I_{WB}(\mathbf{r}) = \left| \int_{-\infty}^{\infty} d\omega I(\mathbf{r}, \omega) \right|^2, \quad (12)$$

where different monochromatic images are added coherently. The same argument applies to TDT and RDT. Moreover, for this reason speckle is present in sonography and DT despite the fact that the insonification is not temporally coherent.

From Eq. (9) it is clear that the speckle phenomenon would not occur if the imaging system were ideal, i.e., if the PSF was a delta function. In this case the image would reproduce the object function exactly. The fact that the PSF is not a delta function is due to the diffraction phenomenon that is the result of interference. This observation explains why CUT is speckle-free. The assumption that $\lambda \rightarrow 0$ implies that diffraction effects are not present and that the PSF is a delta function. In other words, the ray approximation neglects interference that is the very cause for speckle and leads to its total suppression.

Finally it is important to observe that WB DT takes advantage of the entire structure of the wavepackets contained in the time traces recorded by the array for each transmit-receive pair. Each time trace is analyzed in the frequency domain and, at each frequency, the complex number that defines the Fourier transform, at that frequency, is used to produce a cw image. cw images obtained over the bandwidth of the signal are subsequently combined together to produce the final WB DT image. Therefore, if one considers a single time trace, a WB DT image depends on all the complex numbers associated with the Fourier transform of the signal and hence on all the points of the time trace. On the other

hand, for a single transmit-receive pair, CUT uses one real number only that is the delay in the arrival time of the transmitted signal through the specimen relative to the propagation in a background medium. The delay is estimated from particular features of the transmitted wavepacket such as the first zero crossing point or maximum point of the signal envelope. The underlying assumptions are that the wavepacket is rigidly shifted in time by the presence of the object and its amplitude decreased by attenuation while its shape is preserved. Clearly distortion of the wavepacket caused by complex structures within the breast can cause errors in the estimation of the arrival time and result in image artifacts. On the other hand, DT takes advantage of the distortion of the wavepackets to improve image resolution.

II.D. Experimental setup

The experiments are performed with a prototype ultrasound scanner developed at Karmanos Cancer Institute.¹³ The scanner employs a toroidal array with 256 transducers mounted on a circular ring with an internal diameter of 200 mm as shown in Fig. 4(a). The array is immersed in water bath that provides the acoustic coupling between the array and the object to be imaged. The center frequency of the transducers is 1.3 MHz with $\sim 100\%$ bandwidth, further details can be found in Refs. 13 and 32.

The theory above has considered the case of 2D objects that can be reconstructed from measurements performed with a toroidal array. Although the extension of the theory to the 3D case is trivial, limitations in current ultrasound technology, pose fundamental constraints on its practical implementation. In simple terms, while in the 2D case it is sufficient to measure the field with sensors deployed along a circular aperture, in 3D the presence of an additional spatial dimension requires an additional degree of freedom in the measurements. Therefore, the ultrasound field needs to be sampled over a sphere that surrounds the object rather than on a circle. However, in order to avoid aliasing effects, it is necessary to populate the sphere with a number of transducers that approximates the square of the number required in 2D. Even though this condition can be somewhat relaxed, the overall number of sensors that one would need is beyond the capabilities of current technology (for a more detailed discussion, see Ref. 19). To overcome this problem we have introduced a multiscale approach that decouples the information contained in the 2D plane identified by the circular aperture of the array from the global 3D structure of the object, thus allowing a single slice of a 3D object to be reconstructed with a circular aperture.^{19,31} The main idea is illustrated in Fig. 3 and is based on the observation that structures with a characteristic size comparable to λ and contained in the plane of the array can be separated from the global 3D structure of the object using wave refraction. When considering the interaction of the incident field with a 3D object two scales can be identified: (1) the global size of the object; (2) the size of smaller structures contained inside it. In the case of the breast the global size is in order of 100λ , while the structures of interest are in the order of λ or 10λ . Indeed,

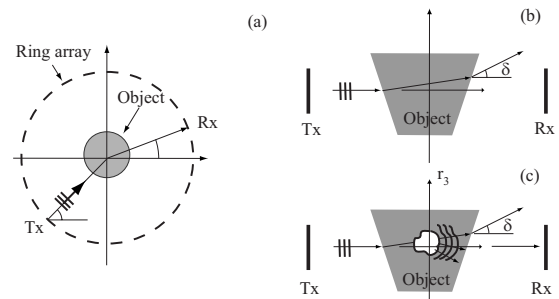


FIG. 3. An illustration of the multiscale method. (a) The array probes a complex 3D object. (b) Cross section of the array with an object showing beam divergence. (c) Forward scattering produced by an internal mass.

λ is chosen so as to ensure that these conditions are met. The wave field emerging from the breast can be divided into two components, a primary field that depends on the global structure of the breast and a weaker secondary field that depends on the smaller structures. The primary field can be described by ray theory and can be separated from the secondary field by using the refraction phenomenon. This causes the primary field to be deflected away from the plane of the array in a similar fashion to the deflection observed when a light beam passes through a prism, Fig. 3(b). On the other hand, the secondary field reaches the array since due to diffraction energy is scattered in almost all directions Fig. 3(c). This process can be optimized by a suitable design of the array parameters, i.e., radius, element size, and frequency, and is demonstrated in Ref. 19.

When a toroidal array is used to detect the secondary field, the sampling requirements are dictated by the wavelength of the incident field and the characteristic size l of the object being probed. In Ref. 33 it has been shown that the minimum number of spatial sampling points N required to avoid aliasing has to satisfy the inequality

$$N > 2\pi l/\lambda. \quad (13)$$

As an example, for an array with $N=256$ and $\lambda=2$ mm the maximum object diameter that can be imaged is ~ 80 mm. For larger objects or shorter wavelengths, aliasing occurs and consequently, it produces grating lobes that interfere with the reconstruction and cause image artifacts.

II.E. Phantom

The phantom, as shown in the photograph of Fig. 4(a), consists of several materials mimicking a subcutaneous fat layer that embeds an irregular glandular region containing four different, 3D inclusions corresponding to two tumors and two fat spheres. The glandular region contains fine powders used to reproduce the speckle effect. However, it does not contain actual structures such as lobes, milk ducts or Copper's ligaments. An X-ray CT image of a coronal slice of the phantom showing all the structures contained in it is displayed in Fig. 4(b), further details can be found in Ref. 13. The phantom was immersed in the water bath at 24.0°C .

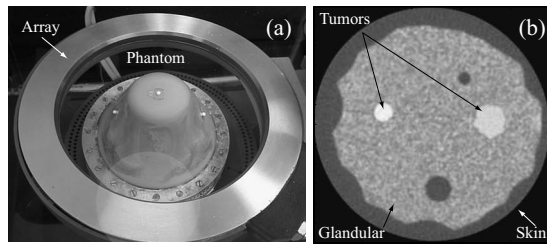


FIG. 4. (a) Photograph showing the complex 3D breast phantom, the toroidal array, and their relative position. (b) X-ray CT of a coronal slice of the phantom showing the four inclusions, the glandular tissue, and the skin.

II.F. Human breast

In order to perform *in vivo* testing, a patient was lying prone on a table with a breast suspended in the water bath through an aperture in the table as described in Ref. 13. The study involved a heterogeneously dense breast that contained an invasive, ductal adenocarcinoma of approximately 25×30 mm in size. The presence of the cancer mass and its size were diagnosed by mammography and conventional B-mode ultrasound; the B-mode image of the breast being shown in Fig. 5. The ultrasound examination showed a poorly differentiated and irregularly shaped hypoechoic mass with associated shadowing. The breast was encircled by the toroidal array in a similar fashion to the phantom experiments and the temperature of the water bath was 22.4°C .

III. RESULTS

This section presents the reconstructions obtained by applying the imaging methods introduced in Sec. II to the ultrasound measurements performed with the prototype scanner for the phantom and the human breast.

III.A. Phantom study

In order to compare different images we define the contrast-to-noise ratio (CNR) as

$$\text{CNR} = \sum_{j=1}^4 \frac{|B_j - B_0|}{\sigma_j + \sigma_0}, \quad (14)$$

where B_j and B_0 refer to the average brightness over the area of the j th mass (the index identifies the four masses in the phantom) and the background, respectively. Similarly, σ_j and σ_0 are the standard deviations of noise calculated by considering the deviation of the point-by-point brightness relative to its mean.

III.A.1. cw images

Figure 6(a) is the sound-speed map of a slice of the phantom obtained with cw TDT at 750 kHz. The image shows the presence and shape of the four inclusions and the skin of the phantom. These characteristics match well with those observed on the X-ray CT image in Fig. 4(b). A careful analysis of Fig. 6(a) also reveals the presence of the irregular contour of the glandular region. However, this is masked by the appearance of the first grating lobe caused by the large diam-

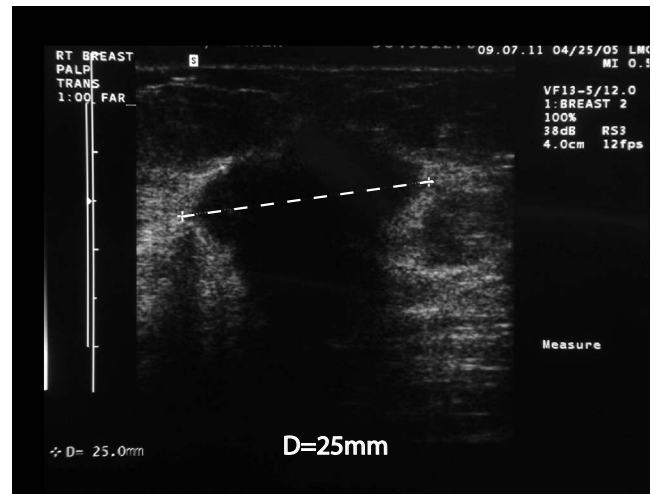


FIG. 5. B-mode image showing a poorly differentiated and irregularly shaped hypoechoic mass with associated shadowing.

eter of the phantom (the phantom diameter is 40 mm larger than the maximum object diameter of 80 mm at $\lambda=2$ mm). The artifact observed between the four inclusions is located at the center of the array and is due to incoherent random noise as shown in the next subsection.

Figure 6(b) is the cw RDT image at 750 kHz. The image is noisy (CNR=0.0) and none of the structures of the phantom can be observed. This is the combined effect of weak backscattering and speckle noise.

Figure 6(c) is the cw RI at 750 kHz. As in the case of Fig. 6(b), the image is noisy (CNR=0.0); however, the outline of the phantom now begins to appear. This modest enhancement over RDT can be explained by observing that RI amplifies the spatial frequencies closer to $2k$, thus improving the reconstruction of boundaries of sudden impedance contrast such as the phantom skin.

III.A.2. WB images

WB images are obtained by coherent superposition of monochromatic reconstructions in the frequency range between 700 and 800 kHz. Note that for the scanner used in this experiments, the lower limit is dictated by the bandwidth of the transducers that have a low output power below 700 kHz, whereas the upper limit is chosen to limit the effects of spatial aliasing since λ decreases as the frequency increases.

Figures 6(d)–6(f) are the WB versions of the corresponding cw images in Figs. 6(a)–6(c). Comparison of Figs. 6(a) and 6(d) shows that WB TDT leads to the suppression of the central artifact, implying that it is caused by random incoherent noise. Moreover, the contrast of the masses relative to the glandular region improves as well as the definition of the glandular region outline. Overall, the reconstruction in Fig. 6(d) exhibits remarkable similarities with the characteristics observed in the X-ray CT image in Fig. 4(b) despite the fact that X-ray CT provides a density map.

The WB RDT image in Fig. 6(e) has an improved CNR = 0.2 compared to the cw RDT of Fig. 6(b); the contour of

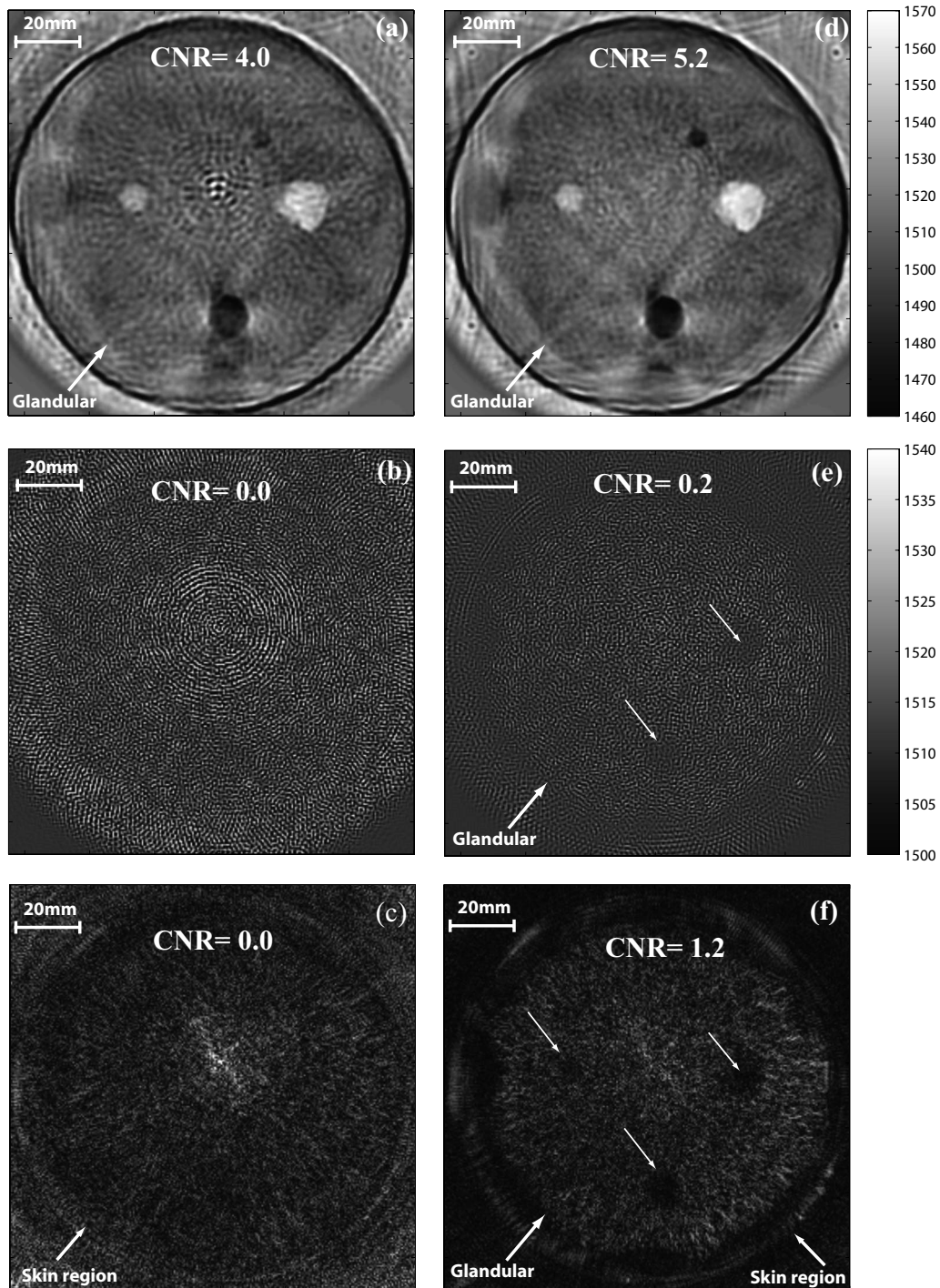


FIG. 6. Coronal slice of the breast phantom reconstructed with different imaging methods: (a)–(c) (left images) cw images at 750 kHz. (d)–(f) (right images) WB images between 700 and 800 kHz. (a) and (d) Transmission diffraction tomography. (b) and (e) Reflection diffraction tomography. (c) and (f) Reflection imaging. The arrows are pointing at the internal masses. CNR values are given at the top of each image.

the glandular region, and two of the inclusions (pointed by the arrows) begin to be visible. However, the estimation of the sound speed is incorrect (the speed inside the inclusions is close to the velocity of the background and for the largest mass this leads to an underestimation of the sound speed by 60 m/s). A greater enhancement is observed for the RI in Fig. 6(f) (CNR=1.2) that improves the reconstruction of the glandular region outline and the skin and shows the third inclusion (pointed by the arrow on the left).

III.A.3. Coherent versus incoherent imaging

Figure 7(a) shows a WB RI over the frequency range between 700 and 800 kHz obtained by the incoherent superposition described by Eq. (11). As pointed out in Sec. II C, incoherent superposition attenuates the speckle contrast, and indeed Fig. 7(a) has a lower contrast (CNR=1.1) than the image obtained with coherent superposition, Fig. 7(b) (CNR=1.2). However, only the reconstruction of the skin of

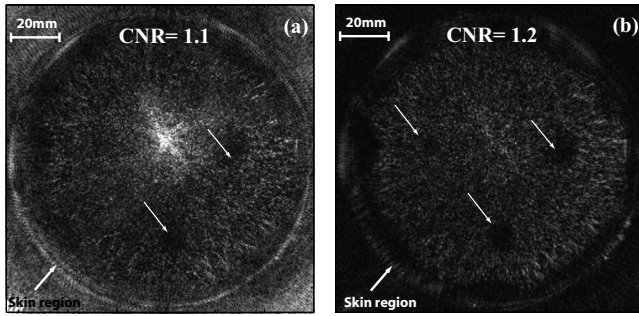


FIG. 7. Speckle suppression by incoherent superposition between 700 and 800 kHz in a reflection image. (a) Incoherent image. (b) Coherent image, same as Fig. 6(f). The arrows are pointing at the internal masses.

the phantom is improved, whereas the image of the interior deteriorates and the small inclusion on the left side is no longer visible. Moreover, the incoherent superposition does not lead to the reduction in incoherent noise and results in a magnification of the artifact at the center of the array.

III.A.4. TDT versus CUT

Figure 8(a) is the CUT sound-speed map of the same slice of the phantom imaged with TDT [Fig. 8(b)]. Despite the fact that CUT uses the full bandwidth of the signal that is centered at 1.3 MHz and the wavelength is approximately one-half of the wavelength used in the TDT reconstruction, the TDT image has superior resolution, providing sharper edges for the masses and more accurate sizing. The CUT image has a higher CNR=7.3 than the TDT (CNR=5.2) image as in the former case noise is lower. However, the contrast relative to the background brightness is higher for the TDT image (4.7) than in the CUT reconstruction (1.4).

On the other hand, the contour of the glandular region is more accurately depicted in the CUT image. This is due to the fact that the aliasing phenomenon that causes the grating lobe in TDT is not present in the CUT image. In fact, under the ray approximation the spacing between the sensors of the

array has to be not larger than the smallest scattering feature in the object to avoid streak artifacts¹⁶ in CUT. This condition is met by the toroidal array for which the ultrasound element spacing is 2.5 mm.

III.B. *In vivo* study

Figure 9 is equivalent to Fig. 6 and refers to a coronal slice of the breast. The images lead to the same observations about cw and WB imaging for the breast phantom; therefore, these will not be repeated here. Instead, attention is paid to the structures contained in the images of Fig. 9. In particular, Figs. 9(a) and 9(d) (the cw and WB TDT images) show the skin of the breast and an entangled network of structures close to the center of the slice. It should be noticed that the breast is larger than the phantom and, therefore, the image artifacts due to the grating lobe are enhanced as confirmed by the blurred reconstruction of the portion of the skin at the top of Figs. 9(a) and 9(d).

The star-point arrows in Fig. 9 point at a region that contains the cancerous mass of approximately 20 mm in size. With reference to Fig. 9(d) it is possible to observe a number of structures departing from the lower part of the mass and penetrating into the surrounding glandular tissue. These same structures result in irregular features along the boundary of the mass that resemble the spiculations observed in B-mode ultrasound images of certain types of malignant tumors.

The mass is not visible in the RTD images Fig. 9(b) and 9(e). However, these show a number of fibrous structures in the lower-right quadrant, close to the skin some of which are pointed by the two arrows. These structures are also found in the transmission image, Fig. 9(d), and the RI image, Fig. 9(f), and could be cross sections of the Cooper's ligaments.

Figure 10 shows a comparison of the CUT reconstruction of the breast with the TDT image as in Fig. 8 for the phantom. However, while in the case of the phantom it is possible to observe a good correlation between the two sound-speed

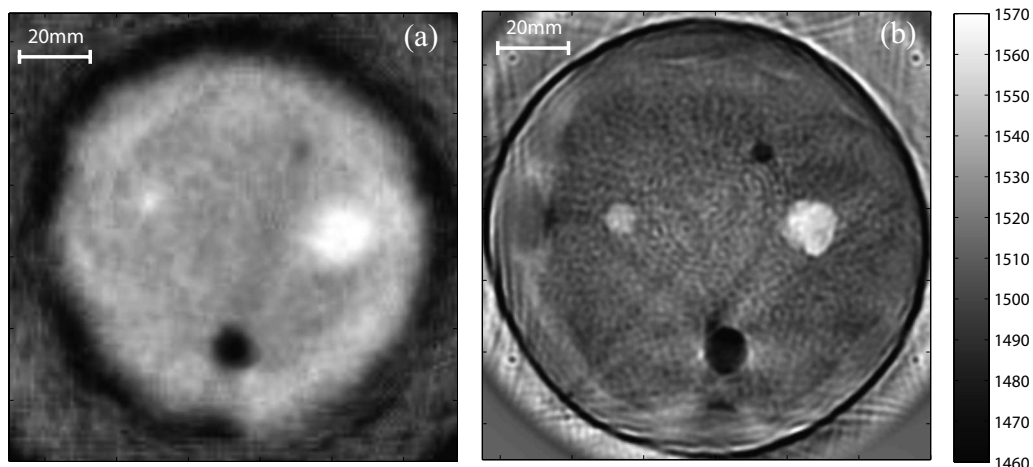


FIG. 8. Transmission tomography images of the phantom: (a) Computerized ultrasound tomography based on the ray approximation. (b) Transmission diffraction tomography, same as Fig. 6(d).

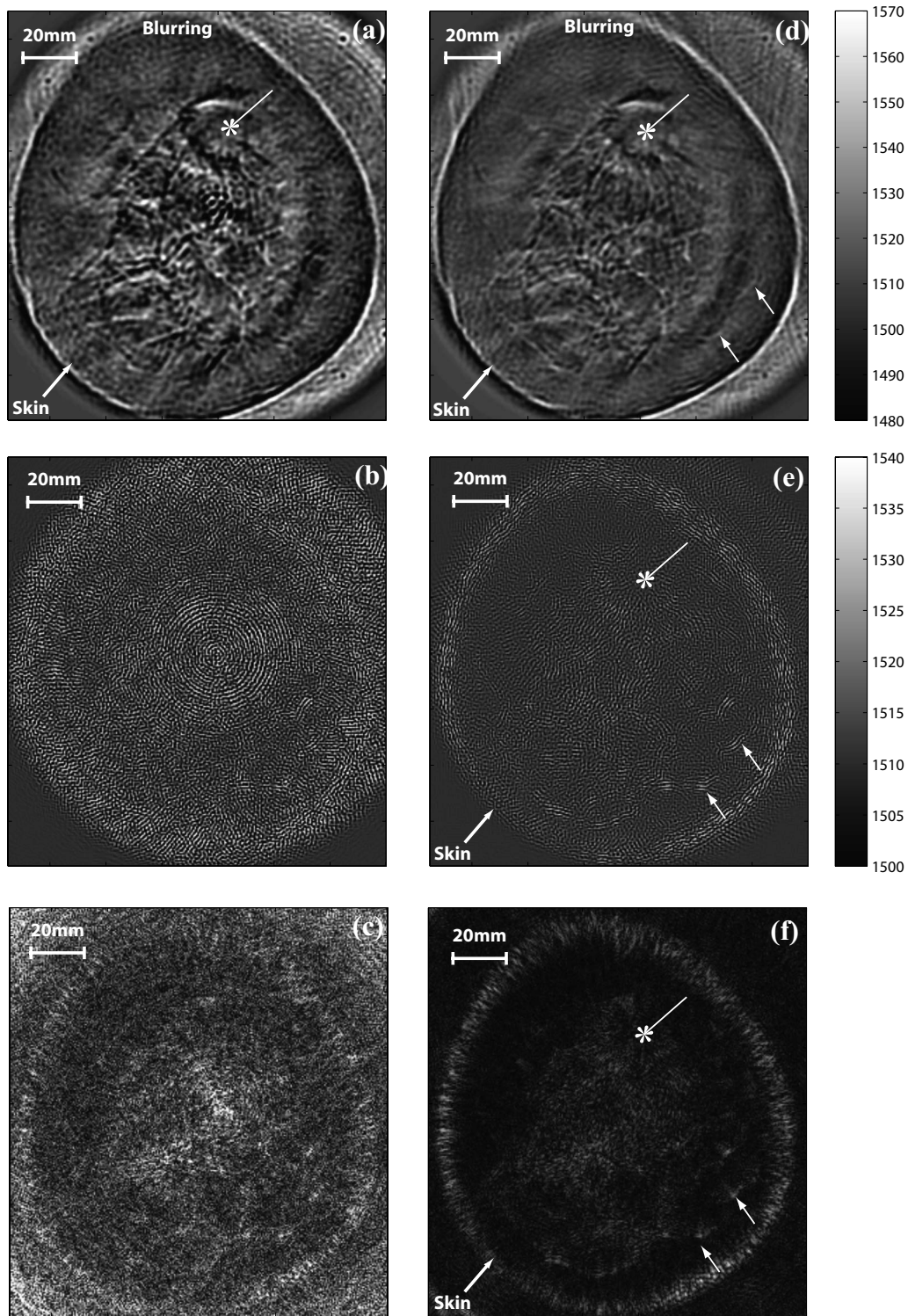


FIG. 9. Coronal slices of a human breast *in vivo*. The star-point arrow points at cancerous mass, while the arrows indicate structures that could be related to Cooper's ligaments: (a)–(c) (Left images) cw images at 750 kHz; (d)–(f) (right images) WB images between 700 and 800 kHz. (a) and (d) Transmission diffraction tomography. (b) and (e) Reflection diffraction tomography. (c) and (f) Reflection imaging.

maps, in the *in vivo* case the same correlation cannot be found. Indeed, the CUT image indicates the presence of a region of high speed of sound at the location of the cancer mass [pointed by the star-point arrow in Fig. 10(a)], whereas the TDT image does not provide a significant deviation from

the sound speed in the glandular region. Moreover, the CUT image shows a number of radial structures that depart from the cancer mass. These appear to radiate at approximately regular angular intervals and correspond to streak artifacts observed in CT images.¹⁶

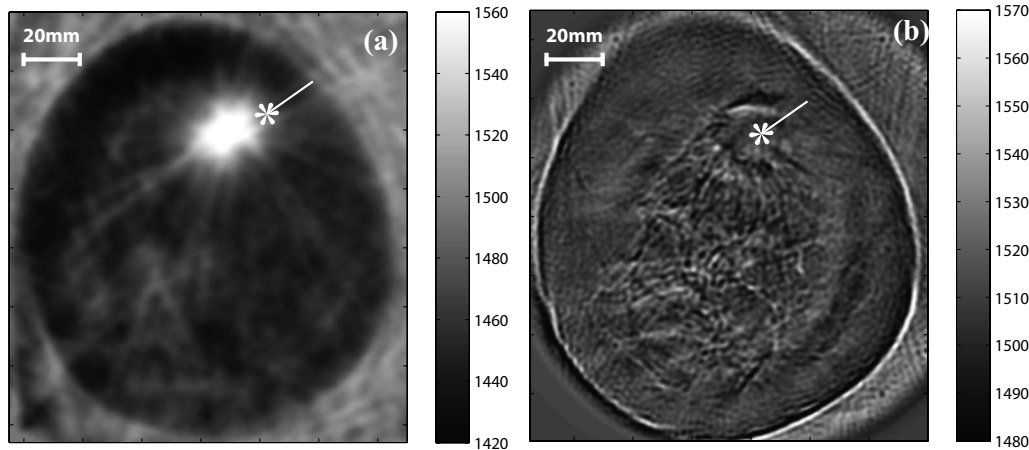


FIG. 10. *In vivo* Transmission tomography images: (a) Computerized ultrasound tomography based on the ray approximation. (b) Transmission diffraction tomography, same as Fig. 9(d).

IV. DISCUSSION

The comparison of the phantom reconstructions obtained with CUT and TDT (Fig. 8) confirms that TDT provides higher resolution than CUT, leading to a more reliable detection and accurate sizing of the phantom inclusions. This demonstrates the importance of diffraction effects in ultrasound tomography. However, when the same comparison is carried out for the *in vivo* data set, CUT indicates the presence of a region of high sound speed at the location of the cancer mass, but that is not observed in the TDT image. The reason for this difference is not clear yet and is the subject of an ongoing investigation.

The network of fine structures observed at the center of the TDT image, which are not visible in the CUT image, could correspond to the milk ducts and Cooper's ligaments. In particular, the fibrous structures visible in the lower-right quadrant of Fig. 10(b), in the subcutaneous fat layer close to the skin (pointed by the two arrows), could be cross sections of the Cooper's ligaments. The same structures can also be found in the WB RDT image, Fig. 9(e), and the WB RI image, Fig. 9(f). Similar structures have recently been observed with the Techniscan system in the paper by Wiskin *et al.*³⁴ In this case the images were obtained by solving the nonlinear inverse scattering problem. Although ductal structures are evident, and cancer is observed little information about its structure is present.

Both the phantom and *in vivo* results confirm that reflection imaging methods complement CUT by providing higher resolution reconstructions of the boundaries of sudden impedance variations. However, this is not true when comparing the reflection images with TDT that exhibits superior image resolution.

Reflection images are much noisier than transmission images. There are two reasons for this. First, the backscattered energy is lower than the transmitted one. As a result, the signal to random incoherent noise ratio is lower in reflection than in transmission. Second, reflection images are based on speckle noise contrast. Speckle is present in reflection images because the scattering produced by the fine structure of

the object contributes to the backscattering measured along the aperture. On the other hand, the forward scattering associated with the same structures is encoded in the ballistic transmission that is deflected away from the object and does not reach the detectors, according to the physical argument proposed in Ref. 19. As a result, transmission measurements are not affected by speckle noise.

Since backscattering measurements are mostly affected by incoherent random noise, WB coherent imaging leads to a greater image noise suppression in reflection than transmission. WB imaging [as described by the functional in Eq. (12)] can lead to reduction of incoherent random noise (not speckle noise) thanks to the averaging effect resulting from the coherent superposition of multiple monochromatic reconstructions. As a result, a wider bandwidth than that used in this paper is desirable because the larger the bandwidth, the larger the ensemble of frequencies over which the averaging occurs. However, as noted in Fig. 7, this does not attenuate speckle noise that can only be reduced by incoherent superposition at the expenses of image signal to incoherent noise ratio.

Finally, the small fat sphere (~ 4 mm diameter) in the breast phantom is not detectable in any of the reflection images as it is masked by speckle noise, whereas it can be clearly observed in the transmission images, indicating that, for the array configuration considered in this paper, transmission measurements are more sensitive to small masses than backscattering measurements. However, it should be stressed that the reflection images shown in this paper refer to a relatively low frequency range (700–800 kHz) to be consistent with the transmission measurements. As a result, the detectability of small lesions with reflection imaging can be improved by increasing the frequency. This is clearly shown in Ref. 13 [Fig. 7(b) in Ref. 13] where by using a wider insonification bandwidth (above 3 MHz) also the smaller inclusion in the phantom becomes visible in the RI.

V. CONCLUSIONS

This paper has investigated the role of diffraction and coherence in breast ultrasound tomography. It has been shown that diffraction effects can be used in transmission diffraction tomography (TDT) to improve image resolution over computerized ultrasound tomography (CUT) that is based on the ray approximation of geometrical optics. For the first time it has been shown that TDT can produce a high-resolution image of a cancer, showing architectural distortion/spiculation as well as the fibrous structure of the breast *in vivo*. Moreover, the *in vivo* study suggests that TDT may be able to image very complex structures inside the breast such as ducts and Cooper's ligaments. However, the same *in vivo* study has shown that while CUT predicts high sound speed inside the cancer mass, TDT leads to the absence of sound-speed contrast between the mass and the surrounding tissue. This potential limitation of TDT is the subject of an ongoing investigation.

While reflection imaging methods complement the information in CUT images, they do not reveal any additional characteristics that are not present in the TDT image at the low frequencies considered in this paper. Indeed, TDT could image a 4 mm inclusion in the phantom that was not visible in the reflection images.

Finally, coherent superposition of continuous wave (cw) images obtained at different frequencies is beneficial to reflection imaging because it reduces the effects of incoherent random noise. The same improvement is not observed in TDT since the signal-to-noise ratio is higher in transmission than in reflection. Therefore, since narrow band transducers have a better response than wideband ones, it can be concluded that narrowband insonification is optimal for TDT.

ACKNOWLEDGMENTS

This work was supported through the U.S. DOE Laboratory Directed Research and Development program at Los Alamos National Laboratory. F.S. is also grateful to the UK Royal Academy of Engineering/Engineering and Physical Sciences Research Council for supporting this work. N.D. and P.L. acknowledge the support of the Michigan Economic Development Corporation (MEDC) under Grant No. MEDC 06-1-P1-0653.

^{a)}Electronic mail: f.simonetti@imperial.ac.uk

¹J. F. Greenleaf, S. A. Johnson, S. L. Lee, G. T. Herman, and E. H. Wood, in *Acoustical Holography*, edited by P. S. Green (Plenum, New York, 1973), Vol. 5, pp. 591–603.

²J. F. Greenleaf, S. A. Johnson, W. F. Samayoa, and F. A. Duck, in *Acoustical Holography*, edited by P. S. Green (Plenum, New York, 1975), Vol. 6, pp. 71–90.

³P. L. Carson, C. R. Meyer, A. L. Scherzinger, and T. V. Oughton, "Breast imaging in coronal planes with simultaneous pulse echo and transmission ultrasound," *Science* **214**, 1141–1143 (1981).

⁴T. M. Kolb, J. Lichy, and J. H. Newhouse, "Comparison of the performance of screening mammography, physical examination, and breast US and evaluation of factors that influence them: An analysis of 27,825 patient evaluations," *Radiology* **225**, 165–175 (2002).

⁵D. T. Ramsay, J. C. Kent, R. A. Hartmann, and P. E. Hartmann, "Anatomy of the lactating human breast redefined with ultrasound imag-

ing," *J. Anat.* **206**, 525–534 (2005).

⁶A. P. Cooper, *On the Anatomy of the Breast* (Longman, London, 1840).

⁷J. G. Abbott and F. L. Thurstone, "Acoustic speckle: theory and experimental analysis," *Ultrason. Imaging* **1**, 303–324 (1979).

⁸J. H. Page, H. P. Schriemer, A. E. Bailey, and D. A. Weitz, "Experimental test of the diffusion approximation for multiply scattered sound," *Phys. Rev. E* **52**, 3106–3114 (1995).

⁹R. L. Weaver and O. L. Lobkis, "Diffuse fields in ultrasonics and seismology," *Geophysics* **71**, S15–S19 (2006).

¹⁰M. P. Andre, M. P. Janeé, H. S., G. P. Otto, B. A. Spivey, and D. A. Palmer, "High-speed data acquisition in a diffraction tomography system employing large-scale toroidal arrays," *Int. J. Imaging Syst. Technol.* **8**, 137–147 (1997).

¹¹F. Lin, A. I. Nachman, and R. C. Waag, "Quantitative imaging using a time-domain eigenfunction method," *J. Acoust. Soc. Am.* **108**, 899–912 (2000).

¹²R. C. Waag, F. Lin, T. K. Varslot, and J. P. Astheimer, "An eigenfunction method for reconstruction of large-scale and high-contrast objects," *IEEE Trans. Ultrason. Ferroelectr. Freq. Control* **54**, 1316–1332 (2007).

¹³N. Duric, L. Littrup, P. Poulo, A. Babkin, R. Pevzner, E. Holsapple, O. Rama, and C. Glide, "Detection of breast cancer with ultrasound tomography: First results with the Computed Ultrasound Risk Evaluation (CURE) prototype," *Med. Phys.* **34**, 773–785 (2007).

¹⁴S. R. Deans, *The Radon Transform and Some of its Applications* (Wiley, New York, 1983).

¹⁵P. R. Williamson and M. H. Worthington, "Resolution limits in ray tomography due to wave behavior: Numerical experiments," *Geophysics* **58**, 727–735 (1993).

¹⁶A. C. Kak and M. Slaney, *Principles of Computerized Tomographic Reconstruction* (IEEE, New York, 1998).

¹⁷A. J. Devaney, "A computer simulation study of diffraction tomography," *IEEE Trans. Biomed. Eng. BME-30*, 377–386 (1983).

¹⁸A. J. Devaney, "A filtered backpropagation algorithm for diffraction tomography," *Ultrason. Imaging* **4**, 336–350 (1982).

¹⁹F. Simonetti, L. Huang, and N. Duric (unpublished).

²⁰J. W. Goodman, *Speckle Phenomena in Optics* (Roberts and Company, Greenwood Village, 2007).

²¹M. Born and E. Wolf, *Principles of Optics* (Cambridge University Press, Cambridge, 1999).

²²M. Azimi and A. C. Kak, "Distortion in diffraction imaging caused by multiple scattering," *IEEE Trans. Med. Imaging* **MI-2**, 176–195 (1983).

²³M. L. Oristaglio, "Accuracy of the Born and Rytov approximations for reflection and refraction at a plane interface," *J. Opt. Soc. Am. A* **2**, 1987–1993 (1985).

²⁴M. I. Sancher and A. D. Varvatsis, "A comparison of the Born and Rytov methods," *Proc. IEEE* **58**, 140–141 (1970).

²⁵M. Slaney, A. C. Kak, and L. E. Larsen, "Limitations of Imaging with first-order diffraction tomography," *IEEE Trans. Microwave Theory Tech. MIT-32*, 860–874 (1984).

²⁶B. Chen and J. J. Stamnes, "Validity of diffraction tomography based on the first Born and the first Rytov approximations," *Appl. Opt.* **37**, 2996–3006 (1998).

²⁷F. Natterer, "An error bound for the Born approximation," *Inverse Probl.* **20**, 447–452 (2004).

²⁸Q. Zhu and B. D. Steinberg, "Large-transducer measurements of wavefront distortion in the female breast," *Ultrason. Imaging* **14**, 276–299 (1992).

²⁹C. W. Manry and S. Broschat, "FDTD Simulations for ultrasound propagation in a 2-D breast model," *Ultrason. Imaging* **18**, 25–34 (1996).

³⁰E. Wolf, "Three-dimensional structure determination of semi-transparent objects from holographic data," *Opt. Commun.* **1**, 153–156 (1969).

³¹F. Simonetti and L. Huang, "From beamforming to diffraction tomography," *J. Appl. Phys.* **103**, 103110 (2008).

³²F. Simonetti, L. Huang, N. Duric, and O. Rama, "Imaging beyond the Born approximation: An experimental investigation with an ultrasonic ring array," *Phys. Rev. E* **76**, 036601 (2007).

³³F. Simonetti, L. Huang, and N. Duric, "On the sampling of the far-field operator with a circular ring array," *J. Appl. Phys.* **101**, 083103 (2007).

³⁴J. Wiskin, D. Borup, S. Johnson, M. Berggren, T. Abbott, and R. Hanover, in *Acoustical Imaging*, edited by M. P. Andre (Springer, New York, 2007), Vol. 28, pp. 183–193.