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## Transparent High-Frequency Ultrasonic Transducer for Photoacoustic Microscopy Application

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C. Yung and J. Hwang are with the Applied Physics Division, National Institute of Standards and Technology, Boulder, CO 80305, USA. Certain commercial equipment, instruments, or materials are identified in this paper in order to specify the experimental procedure adequately. Such identification is not intended to imply recommendation or endorsement by the National Institute of Standards and Technology, nor is it intended to imply that the materials or equipment identified are necessarily the best available for the purpose.

#### Abstract

We report the development of an optically transparent high-frequency ultrasonic transducer using lithium niobate single-crystal and indium-tin-oxide electrodes with up to 90 % optical transmission in the visible to near infrared spectrum. The center frequency of the transducer was at 36.9 MHz with 33.9 %, at -6 dB fractional bandwidth. Photoacoustic imaging capability of the fabricated transducer was also demonstrated by successfully imaging a resolution target and mouse ear vasculatures *in vivo*, which were irradiated by a 532 nm pulse laser transmitted through the transducer.

#### Keywords

Transparent ultrasonic transducer; high-frequency ultrasonic transducer; photoacoustic microscopy

#### I. Introduction

PHOTOACOUSTIC microscopy (PAM) remains the prime choice for label-free microscopic imaging of the optical absorption contrast in tissue [1–3]. It usually employs a single high-frequency ultrasonic transducer to acquire the acoustic waves converted from absorbed optical energy via transient thermoelastic expansion [4–6]. So far, the most widely used PAMs for *in vivo* imaging applications utilize a reflection-mode setup: both the optical illumination and ultrasonic detection placed on the same side of the imaging object [7–11]. However, it imposes certain constraints on PAM's performance. A long working distance is usually required to accommodate the sizeable transducer and the optical-acoustic splitter, thus limiting PAM's numerical aperture and resulting in a poor lateral resolution [12, 13]. Transmission-mode PAMs have achieved finer lateral resolutions, but they are mainly limited to imaging *ex vivo* samples [14, 15]. Further, the complex optical-acoustic splitter, two prisms sandwiching a thin layer of silicon oil, is challenging to make and maintain [12, 13].

To address this problem, doughnut-shaped transducers with a central opening for light passage have been introduced to reduce the obstruction in the optical path [7, 10, 16]. But its size is still too big for PAM to achieve a resolution comparable to optical microscopy [16]. Taking another approach, thin-film transducers that are optically transparent offer a desirable solution. Transparent optic-based ultrasonic detectors include Fabry-Perot etalons [17, 18], micro-ring resonators [19–22], among others [23]. However, they require additional optical instruments such as probe lasers and detectors. Further, it is difficult to integrate these detectors into conventional ultrasound emission devices for ultrasonic/photoacoustic dualmodality imaging. Optically transparent piezoelectric-based devices have been introduced to circumvent this problem. Such devices include ultrasonic transducers with indium-tin-oxide (ITO) coated polyvinylidene fluoride (PVDF) film [24–26], but these PVDF films only achieved a transmittance of 0.8 at best at 532 nm. Also, PVDF's coupling coefficient ( $k_{\rm f}$ ) is not comparable with other popular piezoelectric materials. Another type is capacitive micromachined ultrasonic transducer (CMUT) [27]. This device has limited operating frequency (< 20 MHz) and is of high cost from specialized micro-electro-mechanical systems (MEMS) fabrication techniques.

Moreover, transducers operating at the range of 30-50 MHz are favorable for PAM imaging, since the frequency spectrum of photoacoustic waves after propagating through tissue mainly resides in this range [8]. Therefore, there is a need to find a suitable material for high-frequency ultrasonic transducers with high optical transmission for reflectance mode PAM applications.

Among known piezoelectric materials, lithium niobate (LiNbO<sub>3</sub>) piezoelectric material possesses several properties that are ideal to meet the need. It is a single-crystal material that displays a flat transmittance of 0.8 at a broad range of wavelengths between 350 nm and 5200 nm [28]. It provides good electromechanical coupling, low dielectric constant, and high longitudinal sound speed, ideal for large aperture high sensitivity and high-frequency optically transparent transducers. A brief list of the material properties for LiNbO<sub>3</sub> can be found in [29]. Brodie *et al.* developed an optically transparent piezoelectric transducer with ITO coated LiNbO<sub>3</sub> for ultrasonic particle manipulation [30]. Recently, Dangi *et al.* reported a promising LiNbO<sub>3</sub>-based transparent ultrasonic transducer operating at 14.5 MHz for PA imaging [31].

The purpose of this study is to develop a transducer for PAM applications with a high optical transmission (> 80 %) and a high frequency (> 30 MHz) response. The high transmission of LiNbO<sub>3</sub> single-crystal substrate coated with ITO thin films on both sides is demonstrated over a broad range of wavelengths (450 nm – 1064 nm). PAM imaging performance of the transparent transducer is evaluated by imaging a carbon nanotube-based resolution target and mouse ear vasculatures *in vivo*.

#### II. Materials and methods

#### A. Transducer Design

Parylene thin-film (Parylene C, Specialty Coating Systems, Indianapolis, IN, USA) with nominal acoustic impedance of 2.5 MRayl was selected as the material for the matching layer because it exhibits very little absorption in the visible region and is, therefore, transparent and colorless. To facilitate transducer transparency, an insulating and optically transparent epoxy (EPO-TEK 301, Epoxy Technology, Inc., Billerica, MA, USA) with nominal acoustic impedance of 3.05 MRayl was chosen as the material for backing.

Design parameters of the transducer are summarized in Table I. The inner structure cross section schematic of the transducer is shown in Fig. 1(a). A Krimboltz, Leedom, and Mattaei (KLM) transducer equivalent circuit model-based modeling software (PiezoCAD, Sonic Concepts, Woodinville, WA, USA) was used to simulate and optimize the transducer design [32].

#### **B.** Transducer Fabrication

A 36° rotated Y-cut LiNbO<sub>3</sub> wafer with a dimension of 1 cm  $\times$  1 cm and a thickness of 100  $\mu$ m ( $\lambda$ /2) was acquired from Boston Piezo-optics Inc. (Bellingham, MA, USA) to make a transducer operating at 37 MHz. High optical transmission with surface electrodes was achieved by ITO coatings. Both top and bottom sides of the LiNbO<sub>3</sub> wafer were electroded by sputtering ITO thin films of approximately 108 nm in total thickness. ITO was DC

magnetron sputtered from a 50 mm diameter ITO target (In<sub>2</sub>O<sub>3</sub>/SnO<sub>2</sub> 90/10 wt %, Kurt J. Lesker Company, Jefferson Hills, PA, USA) at 20 W, 933 Pa (7 mTorr, Argon flow  $\approx$  30  $cm^{3}/min$ ) for 80 minutes. The deposition rate was monitored using a quartz crystal microbalance (QCM) with a measured rate of 0.03 nm/s (density =  $7.14 \text{ g/cm}^3$ , z-ratio = 1, tooling = 100 %). For an accurate measure of on-sample deposition rate (LiNbO<sub>3</sub> crystals and QCM are at differing heights), ITO was sputtered on a silicon substrate under similar conditions for 20 min and measured using a white light spectrometer (Filmetrics F40, KLA, Milpitas, CA, USA). The 20 min sputtered film yielded a refractive index of 1.89 and a thickness of 27 nm. Extrapolating these values, we estimate the thickness of the ITO film on the LiNbO3 to be 108 nm. A hook-up wire (Gauge 24, McMaster-Carr Supply Company, Santa Fe Springs, CA, USA) consisting of stranded copper wires was connected to the bottom ITO electrode on the edge of the LiNbO<sub>3</sub> wafer with a small amount of conductive epoxy (E-SOLDER 3022, Von Roll Isola Inc., New Haven, CT, USA). A home-made brass housing with an inner opening of 15 mm in diameter and a subminiature version A (SMA) connector on the side was placed concentrically with the LiNbO3 wafer. The other end of the stranded copper wire was connected to the SMA connector. Degassed epoxy (EPO-TEK 301) with a total weight of 0.2 g was then poured into the brass housing to achieve a 1 mm thick epoxy after curing. For ground connection, a small amount of E-SOLDER 3022 was applied at the four corners on the front surface of the LiNbO3 wafer to connect the top ITO electrode with brass housing. Finally, a 16  $\mu$ m thick ( $\lambda/4$ ) parylene thin-film (Parylene C, Specialty Coating Systems, Indianapolis, IN, USA) layer was vapor-deposited onto the external surface of the transducer by a parylene coater (PDS 2010, Specialty Coating Systems, Indianapolis, IN, USA). A photograph of the fabricated transducer is shown in Fig. 1(b).

### C. PAM Imaging

To demonstrate applications in PAM, we conducted phantom and animal imaging with a reflection-mode PAM [Fig. 2(a)]. The laser beam with 4 kHz repetition rate from a 532 nm picosecond pulsed laser (APL-4000–1064, Attodyne Inc., Toronto, ON, Canada) was focused onto the imaging objects by an aspherical lens (AL2550-A, Thorlabs Inc., Newton, NJ, USA; 50 mm focal length, 0.23 NA), through the transparent transducer. Excited PA waves were detected by the transparent transducer, which was placed in a tank of deionized water for acoustic coupling. A data acquisition unit (ATS9350, AlazarTech, Inc., Montreal, QC, Canada) was used to acquire the PA signal at a 200-MHz sampling rate. The imaging objects were mounted on a two-axis translation stage (PLS-85, PI miCos GmbH, Auburn, MA, USA), and it took ~4 min to acquire an image of 1 mm × 0.8 mm region, with a scanning step size of 3.1  $\mu$ m. Maximum amplitude projection (MAP) images were produced in post-processing by taking the peak-to-peak amplitude of each PA A-line signal.

#### III. Results

#### A. Transducer Performance Evaluation

First, we evaluated the transducer's optical transmission efficiency. An optical parametric oscillator laser (NT242, EKSPLA, Vilnius, Lithuania) provided the illumination at a broad spectrum, and the ratio of the laser light transmitted through the transducer was measured by

a laser power meter (Vega, Ophir Optronics Solutions Ltd., Jerusalem, Israel). Transmission efficiencies greater than 80 % from 450 nm to 1064 nm were measured [Fig. 2(b)]. The trough near 700 nm was also observed on the LiNbO<sub>3</sub>-based transducer in [30].

Electrical impedance and phase angle of the transducer was measured by an impedance analyzer (HP 4294A, Agilent Technologies, Santa Clara, CA, USA) and shown in Fig. 3(b). The resonance frequency ( $f_r$ ) and anti-resonance frequency ( $f_a$ ) of the transducer were measured to be 33.9 MHz and 40.3 MHz, respectively. Using the IEEE standard on piezoelectricity [33], the effective electromechanical coupling coefficient ( $k_{eff}$ ) of the transducer was calculated to be 0.54. It was observed that the measured results are very close to the PiezoCAD simulation results ( $f_r = 31.0$  MHz,  $f_a = 37.8$  MHz, and  $k_{eff} = 0.57$ ) as shown in Fig. 3(a). The electrical impedance of the transducer was measured as 161  $\Omega$  to 258  $\Omega$  around the resonance frequency of the transducer, while the PiezoCAD simulation from 7  $\Omega$  to 22  $\Omega$ . The difference between simulation and measurement especially for electrical impedance could be contributed by the sheet resistance which cannot be simulated in PiezoCAD.

The pulse-echo response of the transducer was measured and analyzed by the method described in [34]. The transducer was mounted on a holder and immersed in a tank filled with deionized water. An X-cut quartz plate was placed about 5 mm away from the transducer as a reflector. The transducer was excited by a pulser-receiver (Panametrics 5900PR, Olympus NDT Inc., Waltham, MA, USA) with 1  $\mu$ J energy per pulse, 200 Hz pulse repetition rate, and 50  $\Omega$  damping factor. The received pulse-echo response and its computed frequency spectrum are shown in Fig. 4(b). The center frequency of the transducer was calculated to be 36.9 MHz, and the –6 dB bandwidth was 33.9 %. Peak-to-peak received voltage ( $V_{p-p}$ ) of the unamplified pulse-echo signal was 0.45 V, indicating the transducer has quite high sensitivity compared to other published transparent transducers [25, 30, 31]. The measured center frequency derived from the pulse-echo result is in good agreement with the PiezoCAD simulation (35.6 MHz) [Fig. 4(a)].

#### B. PAM Imaging with the Transparent Ultrasonic Transducer

Fig. 5(a) shows the MAP image of a resolution target phantom that is composed of carbon nanotube patterns transferred onto a polydimethylsiloxane substrate [35]. The smallest patterns, periodic lines of 10  $\mu$ m width, 100  $\mu$ m long, and 40  $\mu$ m pitch were clearly resolved, while the axial resolution is calculated to be 105  $\mu$ m [36]. The laser pulse energy irradiated on the target was measured to be ~200 nJ, resulting in signal-to-noise ratio (SNR) of ~41.5. Further, we demonstrated the transparent transducer's *in vivo* imaging capability by imaging the mouse ear at the optical wavelength of 532 nm. At this wavelength, the hemoglobin in red blood cells (RBCs) is the primary absorber in tissue, thus enabling imaging the vascular anatomy. The laboratory animal protocols were approved by the Institutional Animal Care and Use Committee of California Institute of Technology. Fig. 5(b) shows the MAP image of a 1 mm × 0.8 mm area of the mouse ear. It can be seen that vessels of a diameter as small as 30  $\mu$ m were clearly imaged.

## **IV.** Conclusions and Discussions

In this work, a transparent high-frequency ultrasonic transducer using ITO coated LiNbO<sub>3</sub> single-crystal was developed for PAM application. The fabricated transducer exhibited an effective electromechanical coupling coefficient  $k_{eff}$  of 0.54, a center frequency of 36.9 MHz, a –6 dB bandwidth of 33.9 %, a high sensitivity of 0.45V peak-to-peak received voltage ( $V_{p-p}$ ), as well as an excellent transparency of up to 90 % in the visible range. Imaging capability of the transducer for PAM was also demonstrated by successfully imaging a resolution target and mouse ear vasculatures *in vivo*, with the laser beam transmitting directly through the transducer. These results will inspire further development of high-frequency ultrasonic transducers in transparent form, which will enable PAM with finer resolution.

Recently, there are quite a few reports on piezoelectric-based transparent ultrasonic transducers fabricated with various techniques [24, 26, 30, 31, 37]. However, this work stands out in the following ways: 1) the transducer's frequency response is the highest, and best matches the frequency spectrum of photoacoustic waves, mainly at the 30 – 50 MHz range, in *in vivo* imaging applications. 2) *in vivo* PAM imaging capability was first demonstrated by this transducer due to its higher sensitivity over others.

Although the transducer demonstrated very good resolution on imaging the resolution target. However, it was not capable to resolve microvessels in the mouse vasculature imaging experiment due to insufficient sensitivity of the transducer. Therefore, there is room for improvement in the performance of transducer. Firstly, the transducer has a relatively high electrical impedance due to high sheet resistance of the ITO thin-film electrodes compared with conventional Au thin-film electrodes [38] which affected transducer sensitivity. It can be improved by choosing a transparent electrode material with lower resistivity. Secondly, choosing suitable materials for both matching layer and backing in ultrasonic transducers can compensate the large acoustic impedance mismatch between piezoelectric element and the load which is a key factor that limits the performance of transducers in terms of sensitivity and bandwidth [39]. Parylene and Epotek-301, both have low acoustic impedance, were chosen as the matching layer material and backing material in the current transducer design, respectively. They limited -6-dB bandwidth of the transducer and degraded transducer sensitivity. Therefore, it is necessary to investigate transparent materials with proper acoustic impedance for the matching layer and the backing in order to improve both sensitivity and bandwidth of the transducer. Furthermore, an optimal front matching scheme incorporating two or more matching layers will also need to be studied.

Employing focusing techniques on the transducer such as using an acoustic lens can also enhance transducer sensitivity. However, focused transducer was not preferred in this work because conventional PAM imaging systems incorporating focused transducers require not only confocal alignment between laser light illumination and acoustic detection, but also mechanical scan of either the transducer or the imaging subjects, which significantly limits PA imaging speed. The next goal of this work is to develop a fast, compact, wearable or hand-holdable PAM imaging device [40], which has broad applications in both *in vivo* biomedical studies and patient monitoring in clinics. A stationary unfocused transparent

transducer, requiring only optical scanning for PAM imaging, has an enormous potential for achieving this goal.

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Biography



**Ruimin Chen** received his B.S. degree in Biomedical Engineering from University of Electronics Science and Technology of China, Chengdu, China in 2006, and both his M.S. and his Ph.D. degrees in Biomedical Engineering from University of Southern California, Los Angeles, CA, in 2008 and 2014, respectively. He worked as a postdoctoral research associate and resource manager in the NIH Ultrasonic Transducer Resource Center at the Department of Biomedical Engineering of University of Southern California. Currently, he is an ultrasound sensor engineer at Masimo Corporation, Irvine, CA.

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Lihong V. Wang is Bren Professor of Medical and Electrical Engineering at Caltech. Published 530 journal articles (h-index = 134, citations = 75,000). Delivered 5350 keynote/ plenary/invited talks. Published the first functional photoacoustic CT, 3D photoacoustic microscopy, and compressed ultrafast photography (world's fastest camera). Served as Editor-in-Chief of the Journal of Biomedical Optics. Received the Goodman Book Award, NIH Director's Pioneer Award, OSA Mees Medal, IEEE Technical Achievement and Biomedical Engineering Awards, SPIE Chance Biomedical Optics Award, IPPA Senior

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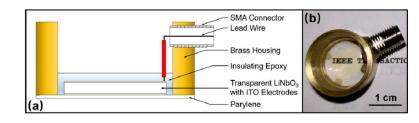
Dr. Zhou is a fellow of the Institute of Electrical and Electronics Engineers (IEEE), the International Society for Optics and Photonics (SPIE), and the American Institute for Medical and Biological Engineering (AIMBE). He is also a member of the Technical Program Committee of the IEEE International Ultrasonics Symposium, and is an Associate Editor of the IEEE Transactions on Ultrasonics, Ferroelectrics, and Frequency Control. His research focuses on the development of piezoelectric high-frequency ultrasonic transducers for biomedical ultrasound and photoacoustic imaging, including intravascular imaging, cancer imaging, and ophthalmic imaging. He is also actively exploring ultrasonic mediated therapeutic technology including trans-sclera drug delivery, as well as ultrasound for retinal and brain stimulation. He has published more than 200 peer-reviewed articles in journals including Nature Medicine, Advanced Materials, and Progress in Materials Science.

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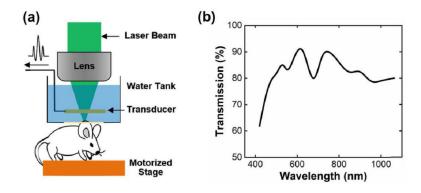
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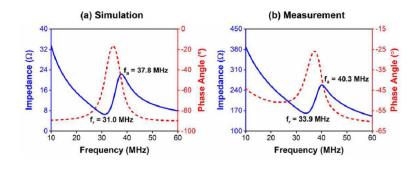


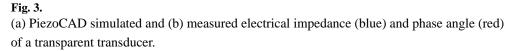
(a) Design cross section of the transparent ultrasonic transducer (not to scale); (b) photograph of a fabricated transparent ultrasonic transducer.

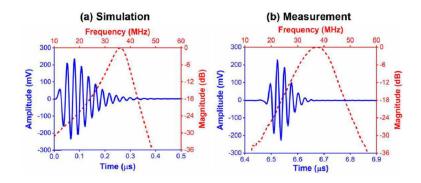


## Fig. 2.

(a) Schematic of the reflection-mode PAM. The transparent transducer, directly in the optical path, is placed on the top of the imaging object to acquire the excited PA waves. The imaging object was mounted on a motorized stage for raster scans; (b) Measurement of optical transmission through the fabricated transducer.

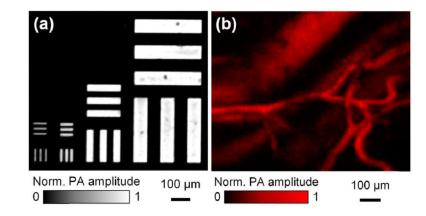






#### Fig. 4.

(a) PiezoCAD simulated and (b) measured pulse-echo response waveform (blue) and its frequency spectrum (red) of a transparent transducer.



#### Fig. 5.

(a) MAP image of the carbon-nanotube pattern. (b) MAP image of an area in the mouse ear showing the vasculatures by exciting the hemoglobin in RBCs. This imaging data was averaged 16 times.

#### TABLE I

#### Design Parameters of the Transparent High-frequency Ultrasonic Transducer.

Specifications	Values
Designed center frequency	37 MHz
Aperture size	$1~{\rm cm}\times 1~{\rm cm}$
Thickness of piezoelectric material (LiNbO <sub>3</sub> )	100 µm
Thickness of matching layer (parylene)	16 µm
Thickness of backing (EPO-TEK 301)	1 mm