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# Lithium niobate transducers for MRI-guided ultrasonic microsurgery

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## Abstract

Focused ultrasound surgery (FUS) is usually based on frequencies below 5 MHz, typically around 1 MHz. Whilst this allows good penetration into tissue, it limits the minimum lesion dimensions that can be achieved. In the study reported here, we investigate devices to allow FUS at much higher frequencies, therefore in principle reducing the minimum lesion dimensions. **Furthermore, FUS can produce deep-sub-millimetre demarcation between viable and necrosed tissue; high frequency devices may allow this to be exploited in superficial applications which may include dermatology, ophthalmology, treatment of the vascular system, and treatment of early dysplasia in epithelial tissue. In this paper** we explain the methodology we have used to build high-frequency high-intensity transducers using Y-36° cut lithium niobate. This material was chosen as its low losses give it the potential to allow very high-frequency operation at harmonics of the fundamental operating frequency. A range of single element transducers with a centre frequency between 6.6 MHz and 20.0 MHz were built and the transducers' efficiency and acoustic power output were measured. A focused 6.6-MHz transducer was built with multiple elements operating together and tested using an ultrasound phantom and MRI scans. It was shown to increase phantom temperature by 32°C in a localised area of 2.5 mm × 3.4 mm in the plane of the MRI scan. Tests on poultry were also performed and shown to create lesions of similar dimensions. This study therefore demonstrates that it is feasible to produce high-frequency transducers capable of high-resolution FUS using lithium niobate.

# Lithium niobate transducers for MRI-guided ultrasonic microsurgery

## I. INTRODUCTION

Focused ultrasound surgery (FUS) is based on the application of high intensity focused ultrasound (HIFU) to heat tissue to a temperature that causes protein denaturation and coagulative necrosis [1]. The required temperature to generate lesions is between 56–60°C [2], [3]. The frequency of ultrasound used is generally around 1 MHz, generating characteristic ellipsoidal lesions on the order of 1 cm in length. Higher frequencies in the region of 4 MHz are also used where more precise treatment is needed, for example in the prostate where tumour sizes may be < 1 mm long [4]. At such frequencies, conventional piezoelectric transducers can be used, based on hard piezoceramic with high drive capability. The use of FUS is increasing as a non-invasive form of surgery and the need for even higher precision is increasing for example for use in aesthetic facial rejuvenation [5], ultrasonic thrombolysis [6] and treatment of malignant disease in breast [7] whilst helping maintain a patient's quality of life when compared to invasive surgery [8].

In this paper, we consider the type of device that could be used to apply HIFU at much higher frequencies, with our research ultimately targeting 50–100 MHz. A difficulty with FUS is to necrose a clinically significant volume quickly enough for financial viability within modern medical treatment systems. However, the exquisite precision of necrosis possible with HIFU must also be recognised. The interface region may be as thin as just a few cells, with deep sub-millimetre dimensions, offering interesting possibilities for precise intervention. These may include treatment of dermatological and ophthalmological conditions, of the walls of the digestive system, for example relating to pre-cancerous tissue dysplasia, and of the walls of the vascular system. The superficial nature of these applications, if necessary handled with intraluminal and intravascular devices, significantly eases the problem of penetration depth at high frequency. The formats of the devices we report here are not suitable for these applications directly but reports on transurethral and endocavitary devices [9], [10] point the way forward. In all cases, we would predict adoption of high frequency FUS for superficial treatments of small overall volumes. As the attenuation coefficient of human tissue has a near linear dependence on frequency [11] greater intensity fields are necessary at higher frequencies in order to be able to penetrate deep enough into human tissue even for superficial applications. Piezoceramic is expected to be incapable of sustaining sufficiently high-power operation at such frequencies because of mechanical fragility, losses, and electrical breakdown. Instead, we have based our investigation on lithium niobate,  $\text{LiNbO}_3$  [12]. As a single crystal, this can be thinned easily without disintegrating

54 unlike ceramics. In addition  $\text{LiNbO}_3$  can sustain high electric fields, and its low losses allow the use of harmonics.  
 55 The use of single crystals and  $\text{LiNbO}_3$  for high frequency ultrasound has been explored before, but only for  
 56 high-resolution imaging [13], [14].

## 57 II. METHODOLOGY

### 58 A. *Lithium niobate*

59 As it was expected that piezoceramics would be unable to produce HIFU at high frequencies and harmonics  
 60 due to de-poling or cracking, we have explored  $\text{Y-36}^\circ$   $\text{LiNbO}_3$ . As well as its basic advantages, it has a high  
 61 resonant frequency of  $3.3 \text{ MHz mm}^{-1}$ , thus allowing for thicker elements at higher frequencies for cost effective  
 62 manufacturing compared to piezoceramics, and it has the highest electromechanical coupling coefficient compared  
 63 to other  $\text{LiNbO}_3$  cuts [15].

### 64 B. *Transducer manufacture*

65 Three transducer designs were prepared as shown in Fig.1: unfocused single elements with 17 mm square  $\text{LiNbO}_3$   
 66 plates (xDucer 1); a 2D faceted bowl with three pentagonal and four hexagonal plates to mimic a spherically-focused  
 67 device (xDucer2); and a 1D faceted cylindrical section with five,  $9 \times 30(\text{mm})^2$  rectangular plates to mimic a  
 68 cylindrically focused device (xDucer3). The equivalent radii of curvature for xDucer2 and xDucer3 were 50 mm  
 69 and 30 mm respectively. **The devices were manufactured as prototypes using a proof-of-concept approach; this**  
 70 **limited their reliability and hence sometimes the completeness of the data that was recorded. Nevertheless, the**  
 71 **manufacturing techniques and the principles of the devices could be taken forward to more robust examples.**

72 To prepare the plates for each transducer,  $\text{Y-36}^\circ$  cut, 3-inch diameter, 0.5-mm thick  $\text{LiNbO}_3$  wafers (Boston  
 73 Piezo-Optics, Inc, Boston, MA) were obtained, polished on one side and lapped on the other. Figure 1 shows the  
 74 position of each element from each wafer for each of the three transducer designs. Separation of the plates was  
 75 performed with a programmable APD1 saw (Logitech Ltd, Glasgow, UK) with a spindle speed of 2900 rpm and a  
 76 feed rate of  $0.160 \text{ mm s}^{-1}$ .

77 For the xDucer 1 devices, the 11 square elements cut from a single wafer were lapped individually in steps of  
 78  $30 \mu\text{m}$  starting from  $500 \mu\text{m}$  down to  $200 \mu\text{m}$  using a PM5 precision lapping and polishing machine (Logitech Ltd,  
 79 Glasgow, UK). The force applied during lapping was adjusted depending on the sample size, typically in the range  
 80  $400\text{--}900 \text{ g}$ . A slurry of  $20\text{-}\mu\text{m}$  calcined  $\text{Al}_2\text{O}_3$  powder in water was used as abrasive. Once the elements reached  
 81 within  $25 \mu\text{m}$  of the target thickness,  $9\text{-}\mu\text{m}$  calcined  $\text{Al}_2\text{O}_3$  powder was used to avoid scratching. The lapping  
 82 machine was programmed to ensure maximum flatness.

83 The true thickness of the samples was measured and verified at regular intervals using a CG-10 Precision  
 84 Electronic Measurement System (Logitech Ltd, Glasgow, UK). Once each element was flat at the desired thickness,

85 it was removed from the glass lapping plate and re-measured to verify the thickness. The elements were continuously  
86 checked using a stereo microscope for flaws which could act to concentrate stress and lead to cracking.

87 An electrode was hand painted on to the lapped side using ELECTRODAG 1415 silver paint (Acheson Colloids  
88 BV, Scheemda, Netherlands). Excess paint around the edges was removed using a scalpel and acetone. The polished  
89 side of each element was then attached to the adhesive side of Adwill D-210 UV tape (Lintec of America, Inc.,  
90 Phoenix, AZ). RG174A/U 50  $\Omega$  coaxial cable was used connected to the plates with Ag-loaded conductive epoxy,  
91 curing taking place at 80°C for 10 min.

92 For xDucer 1 devices, Cu tubing with an internal diameter of 28 mm was cut into lengths of 50 mm and placed  
93 over the LiNbO<sub>3</sub> plates onto the adhesive side of the UV tape. Epoxy was then introduced around the sides of the  
94 LiNbO<sub>3</sub> plate to join it to the Cu tube. The case for the 2D faceted array, xDucer 2, had a height of 75 mm, outer  
95 diameter of 70 mm and a wall thickness of 2 mm. The case for the curvilinear array, xDucer 3, had a height of  
96 75 mm, outer diameter of 50 mm and wall thickness of 1.5 mm. For operation within a magnetic resonance imaging  
97 (MRI) system, the cases of xDucer 2, and 3 were polyvinyl chloride (PVC) coated with a thin layer of Ag paint so  
98 they could be used as the electrical ground connections to the front face of each transducer.

99 To support the fragile LiNbO<sub>3</sub>, Epofix resin (Struers, Ballerup, Denmark) was mixed with S38 glass microballoons  
100 (Lawrence Industries, Tamworth, UK) with a weight ratio of 65 : 35. The microballoon-epoxy mix was poured into  
101 the transducer shell. The xDucer 1 devices were filled to a depth of 16 mm whereas xDucers 2 and 3 were filled to  
102 a depth of 22 mm. It was found that the acoustic output with the backing material was reduced by 5 % compared  
103 to devices made without backing. The backing was left to cure at room temperature. The earth cable was attached  
104 to the shell using conductive Ag epoxy. The UV tape was then exposed to UV light and peeled off. Any remaining  
105 adhesive residue was removed manually.

106 The exposed LiNbO<sub>3</sub> was cleaned using solvent then the front surfaces and part of each case were painted  
107 with Ag paint. The cases were then filled with 5368 silicon (Henkel AG & Co. KGaA, Düsseldorf, Germany) to  
108 waterproof the cables and 50- $\Omega$  BNC RG-174 plugs were connected to the coaxial cables.

### 109 *C. Acoustic pressure*

110 Each transducer was driven by a continuous wave at its fundamental frequency, generated by an AFG3102  
111 waveform generator (Tektronix, Everett, WA). The signal was passed through a -20-dB attenuator before being  
112 used as the input to a power amplifier. The single element transducers were tested using a 3100LA, +55-dB  
113 RF amplifier (Electronics & Innovation, Rochester, NY). To test xDucer 2, the pentagonal elements were linked  
114 and driven by a 2100L, +50-dB RF amplifier (Electronics & Innovation, Rochester, NY) and the hexagonal  
115 elements were linked and driven by the 3100LA amplifier. This was done to give the ability to improve on the

116 alignment of the multiple sound fields by shifting the phase of each group of elements. The pressures outputs  
117 were measured using a calibrated fibre-optic hydrophone (Precision Acoustics, Dorchester, UK) and verified using  
118 an HGL-0200 piezoelectric hydrophone (Onda, Sunnyvale, CA). The curvilinear transducer, (xDucer 3), was tested  
119 using a 150A250, 150W RF amplifier (Amplifier Research, Souderton, PA). The acoustic pressure was measured at  
120 the acoustic focus, 13 mm from the transducer face using the HGL-0200 hydrophone. **The maximum peak-to-peak**  
121 **acoustic pressure was defined at the acoustic focus of each transducer in a  $25 \times 17 \times 12$  (cm)<sup>3</sup>, water filled, low-density**  
122 **polyethylene container. For all measurements the free field was manually scanned to locate the acoustic focus of**  
123 **each transducer using an M-652 *x-y-z* micro-translation stage (Newport, Didcot, Oxfordshire, UK).**

#### 124 *D. LiNbO<sub>3</sub> Properties*

125 Data available for the properties of Y-36° LiNbO<sub>3</sub> were found to be limited and incomplete in the literature so  
126 values for one-dimensional simulation were obtained using PRAP version 2.2 software (TASI Technical Software  
127 Inc, Ontario, Canada) using electrical impedance data from a plate measured with a 4395A impedance analyser  
128 (Agilent, Santa Clara, CA). Table I shows the measured properties for Y-36° cut LiNbO<sub>3</sub>, with figures for Z-cut  
129 material shown for comparison. The resonance frequencies of the transducers were also measured using the same  
130 impedance analyser.

#### 131 *E. Acoustic radiation*

132 The acoustic radiation force output of the transducers was measured using an EMS Model 67 ultrasound radiation  
133 force balance (EMS Physio Ltd, Wantage, UK). The transducers were placed within 20 mm of the surface of the  
134 ultrasound absorber in the balance to ensure that the total radiated flux was incident on it. The output voltage of  
135 the waveform generator was increased and the amplifier forward and reflected power and the transducer acoustic  
136 power were recorded.

#### 137 *F. MRI temperature measurements*

138 MRI guidance is used for FUS [16] as it allows precise targeting of the HIFU field and direct temperature  
139 measurement at the focus. For MRI-guided focused ultrasound surgery (MRgFUS) tests in the present work, xDucer 3  
140 and a DQA Gel Phantom (ATS Laboratories, Bridgeport, CT) placed in a cylindrical perspex chamber filled with  
141 tap water were placed in a GE Signa HDx 1.5T MRI system (GE Healthcare, Waukesha, WI). A gradient echo  
142 planar image (EPI) was recorded with TE = 17.0 ms, ER = 230.0 ms and BW at 62.0 kHz to capture the temperature  
143 increase of the phantom. The curvilinear transducer was turned on at t = 0 s with a pk-pk input voltage of 101 V,  
144 equivalent to 8 W acoustic power and 32 W forward electrical power. The transducer was turned off after 55 s. The  
145 size of the acoustic focus was determined by the area heated above the surrounding ambient temperature.

146 *G. Tissue sonication*

147 To test the effect of the HIFU field on tissue, two boneless, skinless chicken breasts (Tesco, Cheshunt, UK) were  
 148 cut into 12,  $2 \times 2 \times 8$  (cm)<sup>3</sup> strips. The strips were placed in a  $10 \times 15 \times 5$  (cm)<sup>3</sup> container filled with tap water at  
 149 room temperature and xDucer3 was clamped vertically with the acoustic focus on the surface of the tissue. Each  
 150 sample was sonicated once. For tests beneath the surface of the tissue the transducer was lowered closer to the  
 151 chicken. In all experiments it was ensured that the transducer surface was not in contact with the tissue. The chicken  
 152 breast was sonicated using the same settings as in the MRgFUS measurements. Sonication time was increased in  
 153 steps of 10 s. Lesion sizes were measured manually using ImageJ (National Institutes of Health, Bethesda, MD).

154

### III. RESULTS AND DISCUSSION

155 *A. LiNbO<sub>3</sub> Properties*

156 The resonant frequency and third harmonic of the xDucer1 devices made with different LiNbO<sub>3</sub> thicknesses are  
 157 compared to one-dimensional modelling (ODM) for both Z-cut and Y-36° cut LiNbO<sub>3</sub> in Fig. 2. Although the Z-cut  
 158 material gives a higher frequency for a given material thickness, other key properties such as  $d_{33}$  and  $k_T$  are much  
 159 lower, hence the preference for Y-36° cut material.

160 *B. Acoustic pressure*

161 The acoustic pressures generated by the three transducers is given in Table II. The xDucer2 device generated a  
 162 modulated sound field. It was possible to improve the output and reduce the envelope frequency of the modulation  
 163 by shifting the phase of each set of elements. The lowest modulation frequency of 550 kHz was achieved with a  
 164 phase difference of 12°. **The modulated sound field was generated as a result of multiple interacting sound fields.**  
 165 **This was due to misaligned elements in our proof of concept devices. To cancel out the modulation, the phase of**  
 166 **each element would have to be controlled independently, or the faceted bowl must be manufactured to tolerances**  
 167 **at  $< \frac{1}{4}\lambda$  of the operating frequency. Higher harmonics of xDucer2 were not tested due to low reproducibility. The**  
 168 **acoustic pressures generated were limited by the maximum outputs of the RF amplifiers used. From the fundamental**  
 169 **pressure measurements, the design of xDucer2 would be the most logical to pursue.**

170 *C. Acoustic Radiation Power*

171 The acoustic power generated by the xDucer1 devices is shown in Fig. 3. Efficiency for these devices was found  
 172 to be  $33 \pm 5\%$  throughout the frequency spectrum. **Sustained operation of up to 5 minutes was possible without**  
 173 **damaging the transducers at the resonance frequency, 3<sup>rd</sup> and 5<sup>th</sup> harmonic.** The output power is seen to drop as  
 174 the element thickness decreases. This is due to the increasing electrical impedance mismatch shown in Fig. 4. For

175 maximum output power the impedance magnitude and phase should be  $50\Omega$  and  $0^\circ$  respectively. As the element  
 176 thickness decreases the impedance magnitude also decreases. This is seen for both the fundamental resonance and  
 177 3<sup>rd</sup> harmonic. The phase of the 3<sup>rd</sup> harmonic increases with element thickness due to the inductance generated  
 178 by the cable. Factors such as amount of Ag-loaded epoxy and cable length were seen to affect the impedance of  
 179 the transducers [17]. The xDucer3 device had an efficiency of  $25\pm 2\%$  whereas a commercial 3.28-MHz, 58-mm  
 180 diameter HIFU transducer made with piezoceramic (Precision Acoustics, Dorchester, UK) was found to have an  
 181 efficiency of  $20\pm 1\%$ .

#### 182 *D. MRI temperature measurements*

183 Figure 5 is an MRI image of xDucer3 positioned on the DQA gel phantom. Figure 6 shows the area heated by  
 184 xDucer3 in the plane of the MRI scan, aligned with its focus. The surface area of heating after 55 s of sonication  
 185 was  $2.5\text{ mm} \times 3.4\text{ mm}$ . Within 31 s the temperature in the acoustic focus of the transducer had increased  $18^\circ\text{C}$  above  
 186 ambient to a temperature of  $38^\circ\text{C}$ . A peak temperature of  $52^\circ\text{C}$  was reached after 55 s of sonication,  $32^\circ\text{C}$  above  
 187 ambient, as shown in Fig. 7. The acoustic intensity at the focus of the transducer was equivalent to  $163\text{ Wcm}^{-2}$ .  
 188 After sonication, cavitation related bubbles formed on the front surface of the transducer shown in Fig. 5. **These**  
 189 **measurements demonstrate the viability of MRI ultrasonically-elevated temperature measurement with the devices**  
 190 **reported here; it should be noted that at the 3<sup>rd</sup> and 5<sup>th</sup> harmonics (21.1 MHz and 35.2 MHz respectively), the**  
 191 **ultrasound wavelength is substantially smaller than the spatial resolution of the 1.5T MRI system.**

#### 192 *E. Tissue sonication*

193 Figure 8 shows the effect of xDucer3 on chicken breast. An increased sonication time was necessary in order  
 194 to induce a lesion beneath the tissue surface, of similar size to the MRI measured results. This is partially due to  
 195 scattering and gas content in the tissue. The lesions dimensions after 90 s of sonication at 6.6 MHz matched those  
 196 measured with the MRI. In Fig. 8, the chicken has been sliced open to locate the region of sonication. The lesion  
 197 was formed **60 mm** beneath the surface of the tissue without affecting the upper tissue boundary or the surrounding  
 198 tissue. In further tests, after two minutes continuous sonication, when the transducer surface was in contact with  
 199 the chicken tissue, sufficient heat was generated to cause protein denaturation on the chicken surface. **Thus, in a**  
 200 **practical (clinical) setting, contact with tissue would have to be avoided. Impedance matching and better-attached**  
 201 **electrodes might reduce element heating; water-cooling of commercial FUS transducers is also common.**

## 202 IV. CONCLUSIONS

203 In conclusion, we have shown that it is feasible to manufacture high-frequency, high-intensity, focused ultrasound  
 204 transducers based on  $Y-36^\circ$  cut  $\text{LiNbO}_3$ . In a range of tests, we have demonstrated operating frequencies up to

205 more than 50 MHz using the 3<sup>rd</sup> harmonic of 200- $\mu$ m thick LiNbO<sub>3</sub>, focal pressures of 4 MPa at 35 MHz, and  
206 MI = 4.7 at 6.6 MHz. Two of the devices made, with faceted bowl and faceted cylindrical sections respectively, were  
207 designed to be operated under MRI guidance. We have shown that this design was successful and have used one  
208 of the devices to increase the temperature within a gel phantom, measured with MRI, to more than 50°C following  
209 sonication of 55 s with an equivalent acoustic intensity of 163 Wcm<sup>-2</sup>. We also created lesions within chicken tissue  
210 after 90 s sonication.

211 Several aspects can be addressed in order to improve the performance of the transducers. At high acoustic  
212 intensities the Ag-paint electrode was damaged. This is attributed to air pockets trapped between the electrode and  
213 the LiNbO<sub>3</sub>. The use of thin film Cr-Au, Ti-Pt or Al electrodes would be better acoustically and electrically as  
214 electrodes compared to conductive Ag paint [18]. The cases of the devices for MRI guidance were made with PVC  
215 tubing coated with Ag paint; using an alternative such as Cu-epoxy composite [19] would aid manufacture and  
216 reliability and assist with shielding. The thin LiNbO<sub>3</sub> piezoelectric elements were supported by microballoon-filled  
217 epoxy backing; this reduced the transducer output thus necessitating exploration of support materials with a lower  
218 acoustic impedance or other methods to support the plates. Finally, electrical impedance matching was neglected.  
219 However, as frequency increases electrical impedance decreases and sustained operation would be enhanced by  
220 electrical impedance matching.

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1 Transducer manufacturing: (a) position of plates in single wafer used for each transducer, (b) dimensions of plates for each transducer, (c) computer-aided design representation of transducers, and (d) completed transducers. . . . . 11

2 Resonance frequency and third harmonic as a function of element thickness for single-element LiNbO<sub>3</sub> microballoon-backed transducers. The diamonds indicate the z-cut LiNbO<sub>3</sub> simulations, the circles indicate the experimental measurements, and the five-point stars indicate the Y-36° LiNbO<sub>3</sub> cut simulations. The grey data points show the 3<sup>rd</sup> harmonic. The error bars indicate the confidence interval for the LiNbO<sub>3</sub> thickness. . . . . 12

3 Acoustic power as a function of transducer input voltage for 17×17(mm)<sup>2</sup> single element LiNbO<sub>3</sub> transducers. . . . . 13

4 Minimum impedance magnitude and equivalent phase as a function of element thickness for single-element LiNbO<sub>3</sub> microballoon backed transducers. The black diamonds indicate the impedance magnitude whereas the grey circles indicate the phase. . . . . 14

5 (a) MRI side view of sonication setup where brightness indicates water content. Image is rotated 90° anticlockwise from true position. (b) Schematic representation of sonication setup (i) perspex water bath, (ii) xDucer 3, (iii) water, (iv) DQA gel phantom. The transducer generates negligible artefacts in the MRI image. The minor artefacts generated by the silver paint and coaxial cable are not in the acoustic path and thus do not affect the image or temperature measurement. . . . . 15

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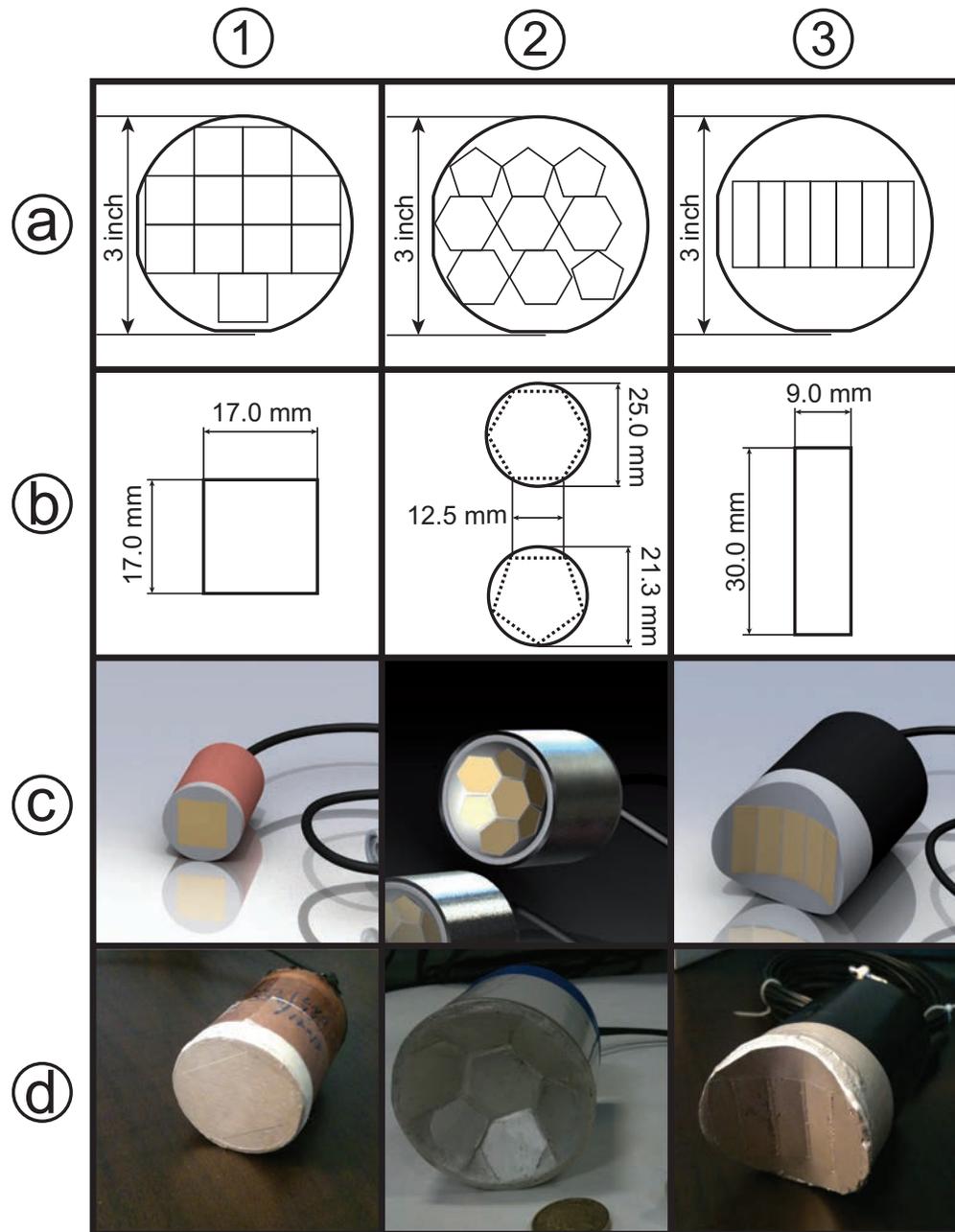


Fig. 1. Transducer manufacturing: (a) position of plates in single wafer used for each transducer, (b) dimensions of plates for each transducer, (c) computer-aided design representation of transducers, and (d) completed transducers.

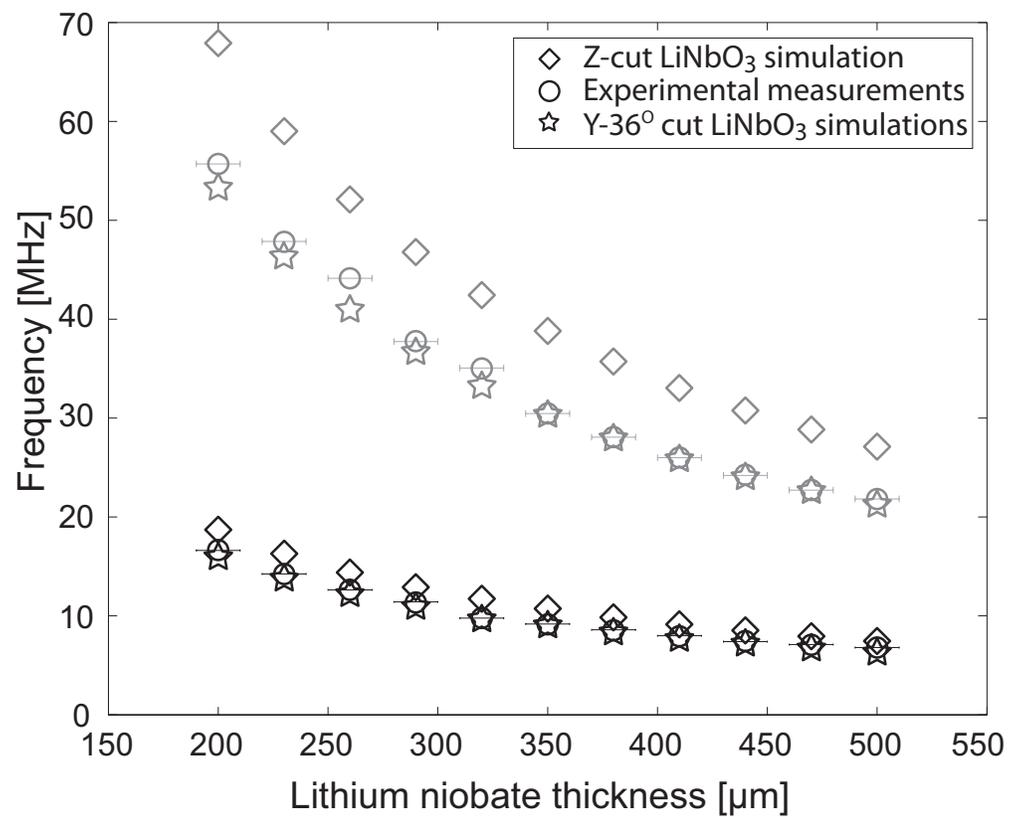


Fig. 2. Resonance frequency and third harmonic as a function of element thickness for single-element LiNbO<sub>3</sub> microballoon-backed transducers. The diamonds indicate the z-cut LiNbO<sub>3</sub> simulations, the circles indicate the experimental measurements, and the five-point stars indicate the Y-36° LiNbO<sub>3</sub> cut simulations. The grey data points show the 3<sup>rd</sup> harmonic. The error bars indicate the confidence interval for the LiNbO<sub>3</sub> thickness.

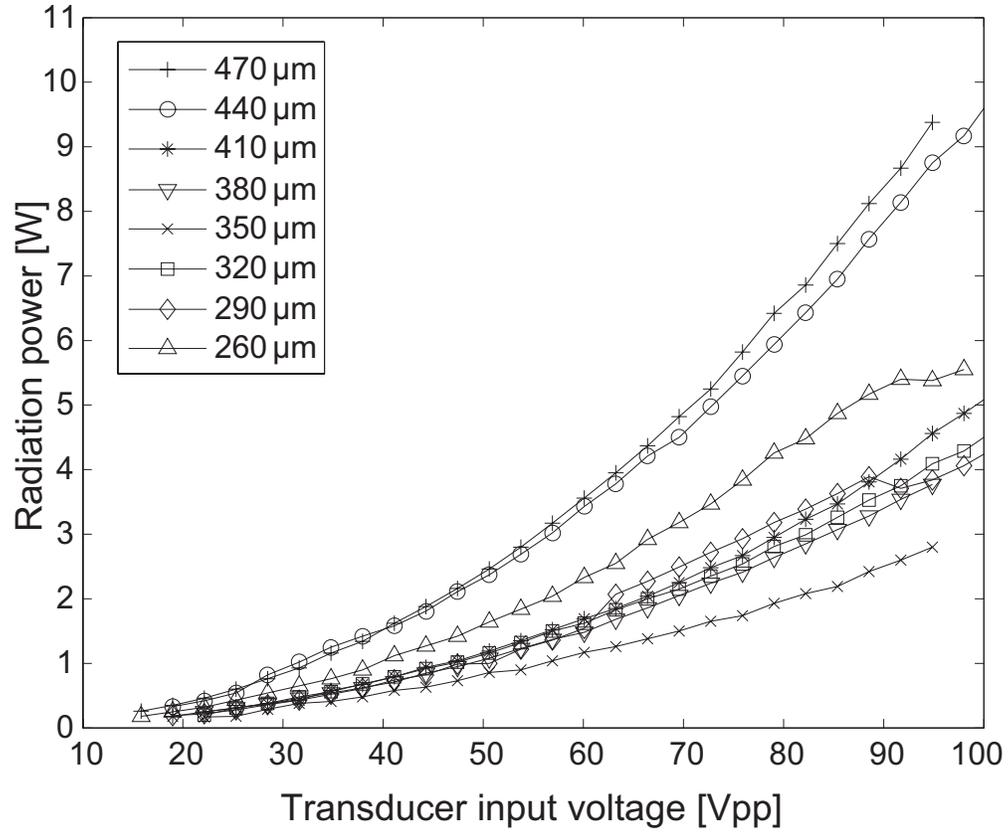


Fig. 3. Acoustic power as a function of transducer input voltage for  $17 \times 17 \text{ (mm)}^2$  single element  $\text{LiNbO}_3$  transducers.

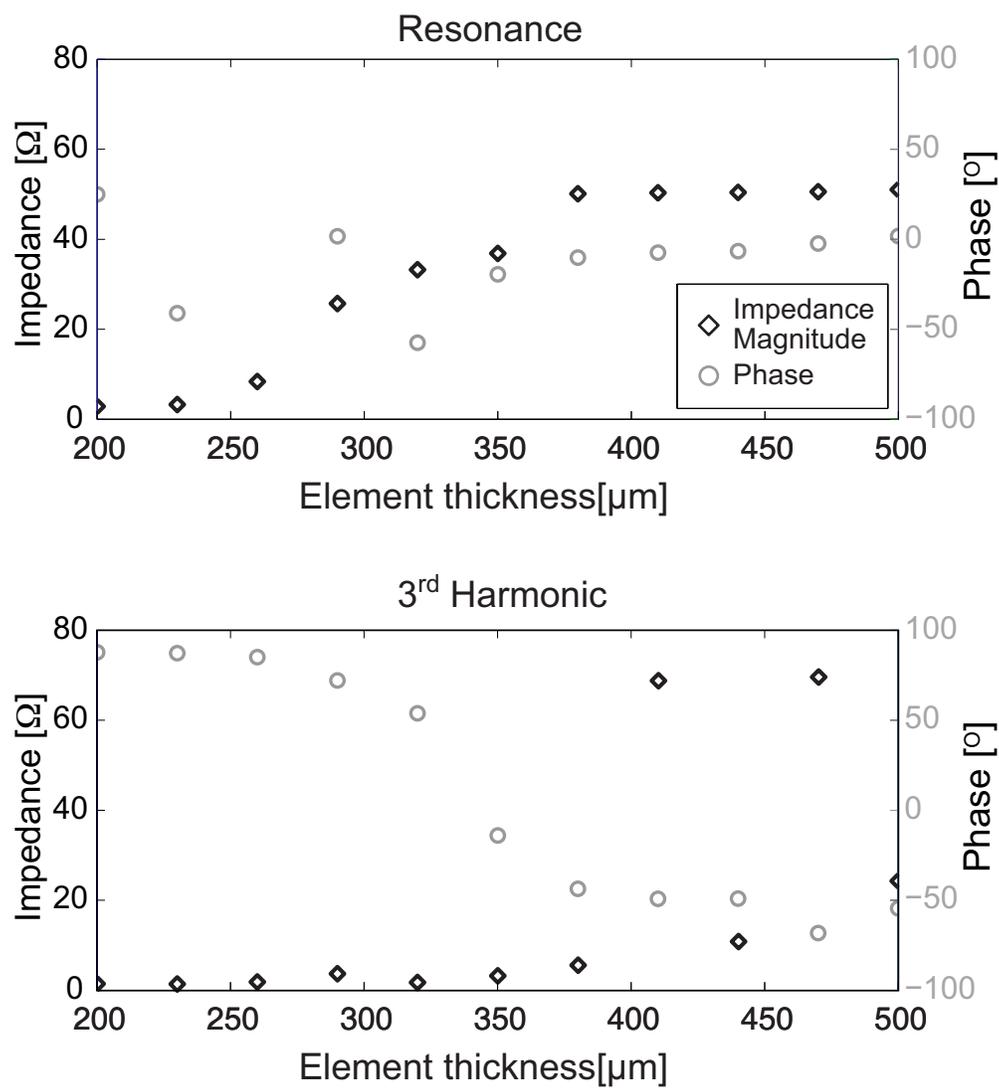


Fig. 4. Minimum impedance magnitude and equivalent phase as a function of element thickness for single-element LiNbO<sub>3</sub> microballoon backed transducers. The black diamonds indicate the impedance magnitude whereas the grey circles indicate the phase.

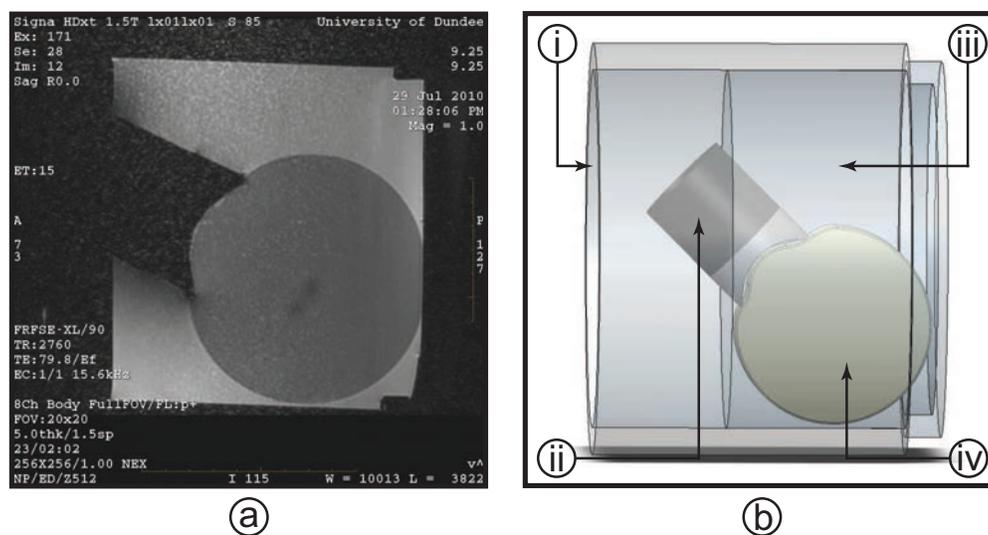


Fig. 5. (a) MRI side view of sonication setup where brightness indicates water content. Image is rotated  $90^\circ$  anticlockwise from true position. (b) Schematic representation of sonication setup (i) perspex water bath, (ii) xDucer 3, (iii) water, (iv) DQA gel phantom. The transducer generates negligible artefacts in the MRI image. The minor artefacts generated by the silver paint and coaxial cable are not in the acoustic path and thus do not affect the image or temperature measurement.

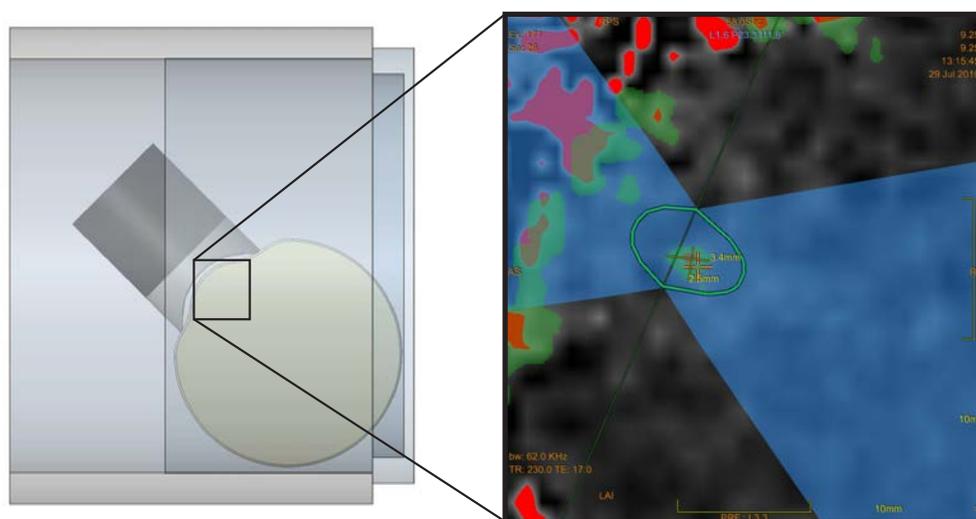


Fig. 6. MRI view of sonication area. The focal region had a size of  $2.5 \times 3.4 \text{ (mm)}^2$ . The green areas represents pixels of equal temperature, the blue areas represent the acoustic field, whereas the red areas represent pixels of temperature  $> 70^\circ \text{C}$ . The orange cross indicates the temperature measurement marker.

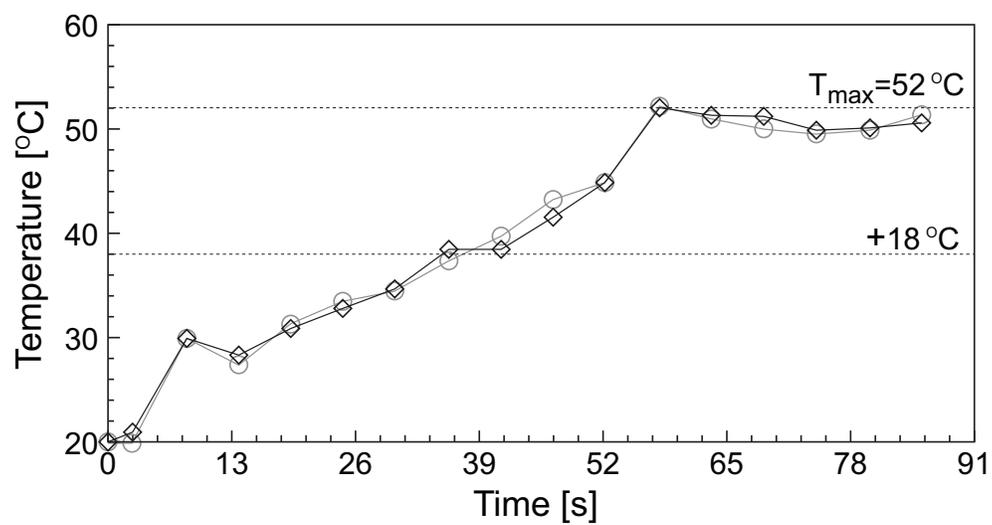


Fig. 7. Temperature increase as a function of time for xDucer3 at the acoustic focus in ultrasonic phantom measured using MRI. The black diamonds indicates the temperature of the selected pixel, whereas the grey circles indicate average temperature of the eight surrounding pixels.

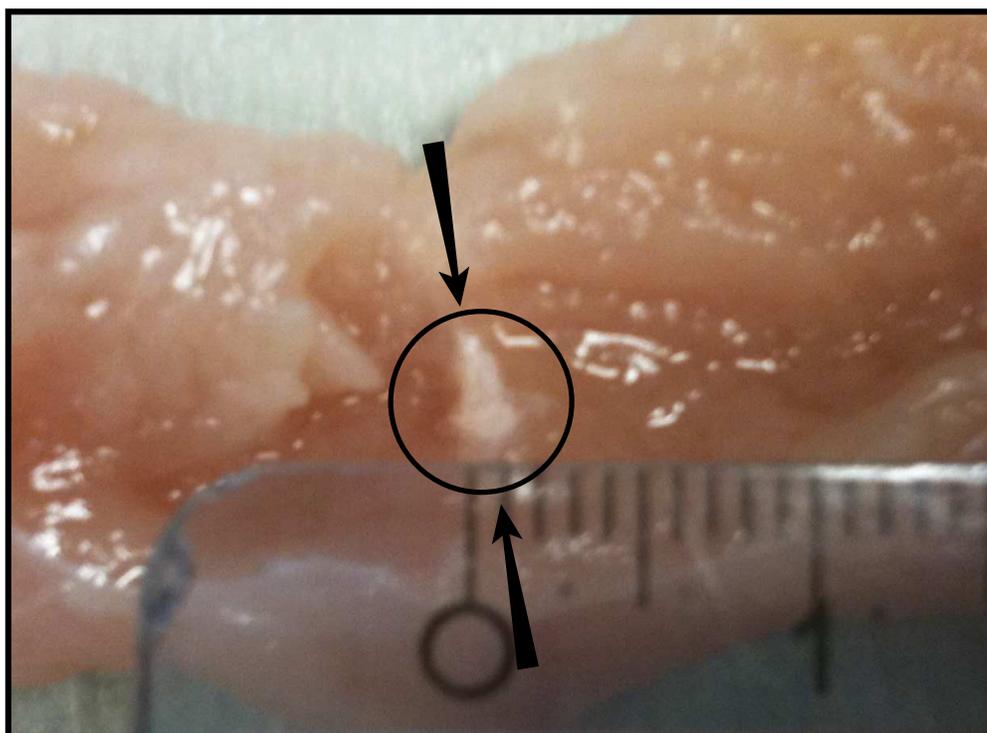


Fig. 8. Photograph showing lesion formed on chicken tissue after 90 s sonication using xDucer 3. The acoustic focus was beneath tissue surface. The direction of acoustic propagation is into the image as indicated by the arrows.

Property	Parameter (unit)	Z-cut	Y-36° cut
Density	$\rho$ (kg m <sup>-3</sup> )	4650	4650
Thickness mode velocity	$v$ (m s <sup>-1</sup> )	7380	7260
Acoustic impedance	$Z$ (MRayl)	34.2	33.8
Elastic constants	$c_{11}^E$ (Nm <sup>-2</sup> ) × 10 <sup>9</sup>	203	185
	$c_{33}^E$ (Nm <sup>-2</sup> ) × 10 <sup>9</sup>	245	185
	$c_{33}^D$ (Nm <sup>-2</sup> ) × 10 <sup>9</sup>	252	245
Dielectric constants	$\epsilon_{33}^T/\epsilon_0$	29.8	41.9
	$\epsilon_{33}^S/\epsilon_0$	25.7	37.6
Piezoelectric constants	$e_{33}$ (C m <sup>-2</sup> )	1.3	4.47
	$h_{33}$ (V m <sup>-1</sup> ) × 10 <sup>9</sup>	5.71	13.4
	$d_{33}$ (m V <sup>-1</sup> ) × 10 <sup>-12</sup>	5.15	18.2
Electromechanical coupling coefficient	$k_T$	0.171	0.495

TABLE I  
MECHANICAL AND PIEZOELECTRIC PROPERTIES FOR LITHIUM NIOBATE

	Frequency (MHz)	Wavelength ( $\mu\text{m}$ )	Peak-to-peak acoustic pressure (MPa) Equivalent mechanical index [MI]		
			xDucer 1	xDucer 2	xDucer 3
Fundamental frequency	6.6	226	14.1 [2.7]	24.3 [4.7]	16.7 [3.3]
3 <sup>rd</sup> harmonic	21.1	77	6.6 [0.7]	-	10.5 [1.1]
5 <sup>th</sup> harmonic	35.2	44	4.3 [0.4]	-	5.4 [0.5]

TABLE II  
PEAK-TO-PEAK ACOUSTIC PRESSURES GENERATED BY xDUCER 1, xDUCER 2 AND xDUCER 3 AT THEIR FUNDAMENTAL FREQUENCY, 3<sup>rd</sup>  
AND 5<sup>th</sup> HARMONIC.

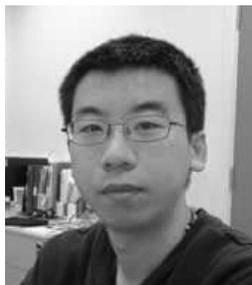
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## VI. BIOSKETCHES



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298 Mechanical Engineering from The University of Hull, England, in 2008. Mr. Kotopoulos is a member of the U.K.  
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302



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309



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317 He has worked extensively with industry internationally and collaborates with several academic research groups.  
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320



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327 the University of Bergen, Norway. Professor Dr. Postema is Associate Editor of Applied Acoustics (Elsevier) and

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329 Institute of Acoustics (IOA), member of the IOA Research Coordination Committee and member of the Scientific  
330 Committee of Revue des Sciences et Technologie (Université de Batna). He has written more than 70 scientific  
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332 particular expertise lies in analysing medical microbubble behaviour under sonication and in high-speed photography.  
333 He also explores non-medical applications of bubbles and droplets in sound fields. Since 2007, he has pulled in  
334 more than US\$4.4 Million in research grants.