

Permanent City Research Online URL: http://openaccess.city.ac.uk/13324/

Copyright & reuse
City University London has developed City Research Online so that its users may access the research outputs of City University London's staff. Copyright © and Moral Rights for this paper are retained by the individual author(s) and/ or other copyright holders. All material in City Research Online is checked for eligibility for copyright before being made available in the live archive. URLs from City Research Online may be freely distributed and linked to from other web pages.

Versions of research
The version in City Research Online may differ from the final published version. Users are advised to check the Permanent City Research Online URL above for the status of the paper.

Enquiries
If you have any enquiries about any aspect of City Research Online, or if you wish to make contact with the author(s) of this paper, please email the team at publications@city.ac.uk.
Heart ablation using a planar rectangular high intensity ultrasound transducer and MRI guidance

Andreas Couppis a,*, Christakis Damianou b,c, Panayiotis Kyriacou a, Cyril Lafon d, Francoise Chavrier d, Jean-Yves Chapelon d, Alain Birerd d

a City University, London, UK
b Frederick University Cyprus, Limassol, Cyprus
c MEDSONIC, LTD, Limassol, Cyprus
d INSERM, U556, Université de Lyon, Lyon F-69003, France

A B S T R A C T

The aim of this study was to evaluate a flat rectangular (3 x 10 mm²) MRI compatible transducer operating at 5 MHz. The main task was to explore the feasibility of creating deep lesions in heart at a depth of at least 15 mm. The size of thermal necrosis in heart tissue was estimated as a function of power and time using a simulation model. The system was then tested in an excised lamb heart. In this study, we were able to create lesions of 15 mm deep with acoustic power of 6 W for an exposure of approximately 1 min. The contrast to noise ratio (CNR) between lesion and heart tissue was evaluated using fast spin echo (FSE). The CNR value was approximately 22 using T1 W FSE. Maximum CNR was achieved with repetition time (TR) between 300 and 800 ms. Using T2W FSE, the corresponding CNR was approximately 13 for the 14 in vivo experiments. The average lesion depth was 11.93 mm with a standard deviation of 0.62 mm. In vivo irradiation conditions were 6 W for 60 s. The size of the lesion in the other two dimensions was close to 3 x 10 mm² (size of the transducer element).

Keywords:
Therapeutics
Hyperthermia
Ultrasound in surgery
Medical ultrasonics

Contents

1. Introduction .................................................................................................................. 821
2. Materials and methods ............................................................................................... 823
   2.1. HIFU/MRI system ................................................................................................... 823
   2.2. Simulation model .................................................................................................... 824
   2.3. Temperature measurement ..................................................................................... 824
   2.4. In vitro experiments ............................................................................................... 824
   2.5. In vivo experiments .................................................................................................. 824
   2.6. MRI processing ....................................................................................................... 824
3. Results ......................................................................................................................... 824
4. Discussion .................................................................................................................... 826
   4.1. Future work ........................................................................................................... 828
References ......................................................................................................................... 828

1. Introduction

Arrhythmia is a problem with the speed or rhythm of the heart-beat. In this condition, the heart can beat too fast, too slow, or with an irregular rhythm. Most arrhythmias are harmless, but some can be serious or life threatening. With arrhythmia, the heart may not be able to pump enough blood to the body [1] Lack of blood flow can damage the brain, heart, and other organs. Atrial fibrillation (AF) and Ventricular Fibrillation (VF) are the most common types of serious arrhythmias.

AF is a very fast and irregular contraction of the atria. AF occurs when the heart's electrical signal begins in a different part of the atrium than the sinoatrial (SA) node or when the signal is conducted abnormally [1–3].
VF occurs when disorganized electrical signals make the ventricles quiver instead of pump normally. Without the ventricles pumping blood out to the body, a person will lose consciousness within seconds and will die within minutes if not treated [1]. To prevent death, the condition must be treated immediately with defibrillation, an electric shock to the heart.

The Cox–Maze procedure is the gold standard for the surgical treatment of atrial fibrillation (AF) [4–7]. Despite its efficacy, this procedure is not widely performed because of its complexity and technical difficulty. Recently, the introduction of ablation technologies has significantly changed this attitude. To simplify the operation, the incisions of the traditional cut and sew Maze procedure have been replaced with linear lines of ablation.

Ablation technologies have greatly simplified surgical approaches and have significantly increased the number of procedures being performed [8]. Various energy sources have been used clinically, including cryoablation, radiofrequency, microwave, laser, and high-frequency ultrasound. The goal of these devices is to create lesions to block activation wavefronts [8].

Cryoablation is unique among presently available technologies in that it destroys tissue by freezing instead of heating. The biggest advantage of this technology is its ability to preserve tissue architecture and collagen structure [9]. The potential disadvantage of this technology is the relatively long time necessary to create an ablation lesion (2–3 min). Also, the circulating blood volume acts as a heat sink which makes creating lesions in the beating heart difficult [9].

In electrophysiology laboratories, RF energy has been used for cardiac ablation for many years [10]. It is one of the first energy sources to be used in the operating room for AF ablation. The first team to use RF energy in humans was that of Borggreve et al. [11] who disconnected a right-sided accessory pathway. Their pioneering work, which was subsequently followed by many other teams, opened the door to transvenous ablation at a significantly lower risk compared with direct current (DC) ablation. This rendered RF ablation a widely applicable therapy for patients with supraventricular and ventricular arrhythmias. The efficacy of RF ablation is limited in cases which require the ablation of larger areas in thicker tissues, and when ablation is performed on the left side of the heart the risk of charring and thrombus formation becomes an issue, especially when high power settings and high target temperatures are used. A major advance in this area came after a long series of animal experiments mainly performed by Wittkampf [12] using irrigated tip RF ablation with closed or open irrigation.

Microwave ablation is an interesting technology that has several advantages over RF [13]. It is less likely to create char formation and is less sensitive to electrode positioning. It can reliably create transmural endocardial lesions, but it is not capable of creating epicardial lesions on the beating heart. Microwave ablation is an unfocused energy source that can cause collateral injury. This has been documented in case reports [13].

Laser ablation is a promising technology that may have some advantages over other energy sources [14–16]. The energy is focused, unaffected by overlying fat, and uses flexible fiber optics to deliver the energy to tissue. A disadvantage of this technology is that the energy delivery is unconfined and thus could cause collateral damage.

Therapeutic ultrasound, is a non-invasive extracorporeal technique capable of thermally ablating subsurface structures without injuring intervening tissues. Ultrasonic energy can be applied in a target volume to induce molecular agitation, absorptive heating, and ultimately thermal coagulative tissue necrosis. Therapeutic ultrasound offers several potential advantages over other therapy modalities. Unlike microwaves, it can be readily focused within small volumes. It does not have the cumulative risks associated with ionizing radiation, and it is unaffected by the optical opacities that block laser penetration. Furthermore, in contrast to radiofrequency (RF) waves, therapeutic ultrasound does not require direct contact with target tissue. The use of therapeutic ultrasound to create focal, ablative lesions in the heart has been reported in experimental models [17–19].

Therapeutic ultrasound was explored as a therapeutic modality in almost every accessible tissue. Several studies have examined the histological changes related to therapeutic ultrasound ablation in various tissues [20,19,21,22,18,23–27].

Within the last decade, therapeutic ultrasound has been used in various in vitro models [28–32]. Okumura et al. in 2008 [31], performed a study to examine tissue temperatures around pulmonary veins (PVs) during therapeutic ultrasound balloon ablation for atrial fibrillation. In 2009, Yokoyama and colleagues [30] developed a canine model of esophageal injury and left atrial-esophageal fistula after applications of forward-firing therapeutic ultrasound and side-firing unfocused ultrasound. Neven et al. in 2010 [29], evaluated an esophageal temperature (ET)-guided safety algorithm to apply therapeutic ultrasound in order to avoid severe complications with standard ablation of pulmonary vein isolation (PVI).

Therapeutic ultrasound has been used in various in vivo models for heart applications: Strickberger et al. in 1999 [18], carried out a study to determine whether therapeutic ultrasound can be used to ablate the atroventricular (AV) junction within the beating heart. Their study suggested that therapeutic ultrasound produced well-demarcated lesions and appears to be a feasible energy source to create complete AV block within the beating heart without damaging the overlying or underlying cardiac tissue. Otsuka et al. [33] investigated the possibility of myocardial ablation in the left ventricle of beating dog hearts by monitoring treatment with 2-dimensional echocardiography. Recently a transesophageal ultrasound applicator was designed and evaluated with in vivo experiments [34,35] for the treatment of atrial fibrillation. The results indicated a potential application of the transesophageal ultrasound applicator for atrial fibrillation treatment.

Over the last decade, clinical studies have been conducted using commercially therapeutic ultrasound systems. Early study [36], revealed that, therapeutic ultrasound can create precise defects in different cardiac tissue without damage to the surrounding tissue. A multicenter clinical trial in 2005 [37], showed that after 6-month follow-up, epicardial, off-pump, beating-heart ablation with acoustic energy is safe and cures 80% of patients with permanent atrial fibrillation associated with long-standing structural heart disease.

A therapeutic ultrasound balloon catheter to isolate pulmonary veins (PVs) outside the ostia (PV antrum) was designed by Nakagawa et al. in 2007 [38]. Results of this study showed that forward-focused therapeutic ultrasound application isolated PVs outside the PV ostium with elimination of AF in 16 (59%) of the 27 patients at 12 months following the single ablation procedure. Another study by Schmidt et al. in 2009 [39] showed that, the novel defocused 12F Therapeutic Ultrasound-Ballon Catheter used in conjunction with a steerable sheath allows for very rapid pulmonary vein isolation (PVI) in patients with paroxysmal atrial fibrillation (PAF). Epicard cardiac ablation system [40,41] has been proven to be a concomitant procedure during open heart surgery that is safe and acceptably effective for the treatment of AF. Metzner et al. in 2010 [42], performed a clinical study for pulmonary vein isolation using first and second generation therapeutic ultrasound balloon catheters (ProRhythm, Ronkonkoma, NY, USA). Patients treated with symptomatic PAF showed long-term success rates similar radiofrequency current (RFC) ablation-based PVI procedures.

The combination of ultrasound and MRI was first cited by Jolez and Jakab [43] who demonstrated that an ultrasonic transducer can be used inside an MRI scanner. The concept of using MRI to monitor the necrosis produced by therapeutic ultrasound was demonstrated in the early 90's by Hynynen et al. [44] in canine
muscle. In the following years many studies have been conducted showing that the contrast between necrotic tissue and normal tissue was excellent.

In this paper a flat rectangular MRI compatible transducer, $3 \times 10 \text{ mm}^2$, operating at a frequency of 5.3 MHz was used. The aim was to create lesions in lamb heart in vitro and in rabbit in vivo using the cardiac transducer in order to measure the size of thermal necrosis in relation to the power and time. The experimental results are compared with the results of a simulation model.

Another task was to investigate the effectiveness of MRI in monitoring therapeutic protocols of therapeutic ultrasound in the heart. This was achieved with the optimization of MRI pulse sequences in order to maximize contrast between thermal necrosis and healthy cardiac tissue. Screening with MRI makes therapeutic ultrasound a more credible way of treatment in comparison to other ablation methods which are guided by X-rays. MRI offers a better contrast between thermal necrosis and healthy cardiac tissues, and offers the additional advantage that the patient will not be exposed to radiation. Therefore we have used the basic MRI sequences T1-W and T2-W fast spin echo (FSE). With T1-W FSE, the effect of repetition time (TR) on contrast was explored, whereas with T2-W FSE, the effect of echo time (TE) was explored.

The main innovation of this study is the evaluation of a planar MR compatible transducer in both in vitro and in vivo exploring the following 2 main issues:

(a) Acquire knowledge regarding the size of the lesion as a function of power and time (using simulation and experimental models) which could be used as a preliminary data for future clinical trials and

(b) Optimize the FSE MRI techniques for detecting the lesions created in heart tissue.

The advantage of using MRI is that the lesions can be monitored nearly in real time and therefore unwanted complications similar to what was reported by Metzner et al. in 2010 can be avoided.

2. Materials and methods

2.1. HIFU/MRI system

Fig. 1 shows the block diagram of the therapeutic ultrasound/MRI system. The therapeutic ultrasound system consists of a signal generator (HP 33120A, Agilent technologies, Englewood, CO, USA), a RF amplifier (250 W, AR, Souderton, PA, USA), and a flat rectangular MR compatible transducer (Fig. 2).

The transducer consisted of a 3.8-mm outer diameter copper tube with a 0.1-mm thick wall ending in a cone-shaped MR compatible plastic tip. The plane active surface embedded in the plastic section is a $3 \times 10 \text{ mm}^2$ P762-type PZT piezoceramic air-backed transducer (Quartz & Silice, Nemours, France) operating at 5.3 MHz. In order to heat the myocardium for the purpose of treating arrhythmias, the transducer must be placed in a catheter. Because the catheter is inserted into the body, the catheter is destroyed after treatment (i.e. it is considered as consumable). Thus, a spherically focused technology has the additional disadvantage that will increase the cost substantially. Additionally, as it will be seen from the results, a flat transducer produces lesions of desired size.

The central lumen of the tube provided electrical radiofrequency (RF) connections and a path for the cooling water. RF connections were made via a miniaturized 50-cm long, 50 $\Omega$ coaxial cable with a 0.9-mm outer diameter. The outer ground conductor was connected to the external face of the transducer. The inner conductor reached the internal face of the transducer through the lumen of the tube. The transducer was glued with an MR compatible epoxy resin (Emerson & Cuming Europe nv, Westerlo-Oevel, Belgium) to ensure that it was watertight and to hold it in place. To limit energy loss during transmission of the electric signal, a connection with a capacitor-inductor network was included to match the transducer. Using the acoustic balance technique by
Davidson 1991 [52], the electroacoustic efficiency of the applicator was measured to 65% at 5.3 MHz. The external face of the transducer was cooled by a continuous flow of degassed water circulating the length of the transducer. The cooling water serves as coupling medium between the transducer and the heated tissues. The water cooling circuit was maintained at 15 °C and was driven by a Masterflex peristaltic pump (Cole Parmer Instrument Co., Chicago, IL) at a flow of 0.15 L/min.

2.2. Simulation model

Numerical simulations were performed to predict the shape of the thermally ablated zone for different strategies, varying the following input parameters: thickness of the cooling water, the elementary sonication duration and the acoustical power. The model was proposed by Chavrier et al. 2000 [53] improved later by Curiel et al. 2004 [54]. It is based on the Bio Heat Transfer Equation proposed by Pennes [55]. Simulations neglected cavitation as the evaluated transducer is not focused and operates in the high frequency range.

The initial temperature of the heart was assumed to be 37 °C and the initial temperature of the water was assumed to be 15 °C. The spatial step used during the simulation was 0.5 mm and the temporal step used during the simulation was 0.1 s. The thermal dose threshold of necrosis was considered to be 240 min referenced at 43 °C.

The numerical model was implemented in C++ supported by a graphic user interface developed in MATLAB (MathWorks, Natick, United States). The simulation environment provided the following output: a 3D view plus two central sections of the predicted ablation zone, total volume of the coagulation necrosis and 3D map of estimated thermal dose.

Numerical simulations were performed for some exposures in vivo conditions (the tissue parameters used are shown in Table 1).

2.3. Temperature measurement

Temperature was measured using a Digital to analog data acquisition card (USB 6250, National Instruments, Austin Texas, USA) and a voltage to temperature converter (Omega Engineering, Stamford, Connecticut, USA). Temperature is sensed using a 50-μm diameter T-type copper-constantan thermocouple (Physitemp Instruments, Inc. New Jersey, USA) which is MRI compatible. The thermocouple was placed between the tissue and the transducer surface, thus measuring the temperature at the surface of the tissue. The temperature error of the thermocouple was on the order of 0.1 °C. In all the experiments, the goal was to keep the temperature below 100 °C in order to avoid boiling.

2.4. In vitro experiments

Various in vitro experiments were carried out to image the lesions using MRI created in cardiac tissue using therapeutic ultrasound. The tissue was placed inside the water container which was filled with degassed water at room temperature. The transducer was coupled to the holder and was immersed in the water tank, thus providing good acoustical coupling between tissue and transducer (see Fig. 1 for the experimental arrangement). In all experiments, the tissues used were extracted from freshly excised lamb, and the experiment was conducted in the same day. In total, 18 samples were used.

2.5. In vivo experiments

For the in vivo experiments, adult rabbits from Cyprus were used weighting approximately 3–4 kg. 14 rabbits in total were used in the experiments. The rabbits were anaesthetized using a mixture of 500 mg of Ketamine (100 mg/mL, Aveco, Ford Dodge, IA), 160 mg of Xylazine (20 mg/mL, Loyd Laboratories, Shenedoah, IA), and 20 mg of Acepromazine (10 mg/mL, Aveco, Ford Dodge, IA) at a dose of 1 mL/kg. The protocol for the animal experiments was approved by the national body in Cyprus responsible for animal studies (Ministry of Agriculture, Animal Services).

2.6. MRI processing

An MRI scanner (Signa 1.5 T, by General Electric, Fairfield, CT, USA) was used. The ultrasound transducer is coupled to the holder. The spiral coil (USA instruments, Cleveland, OH, USA) was used to acquire the MR.

The following parameters were used for T1-W FSE: TR was variable from 100 to 1000 ms, TE = 9 ms, slice thickness = 3 mm (gap 0.3 mm), matrix = 256 × 256, FOV = 16 cm, NEX = 1, and ETL = 8. For T2-W FSE: TR = 2500 ms, TE was variable from 10 ms to 160 ms, slice thickness = 3 mm (gap 0.3 mm), matrix = 256 × 256, FOV = 16 cm, NEX = 1, and ETL = 8.

The CNR was obtained by dividing the signal intensity difference between the Region of Interest (ROI) in the lesion and in the ROI of normal tissue by the standard deviation of the noise in the ROI of normal tissue. The ROI was circular with diameter of 3 mm.

3. Results

Fig. 3 shows T1 W FSE images of the 2 transducers utilized during the experiments in sagittal (Fig. 3A) and coronal (Fig. 3B) planes. Note the RF artefacts of the transducers are minimal. There is a 1 mm artefact around the transducer due to the copper material of the transducer and of the piezoelectric element.

Fig. 4A shows the lesion depth (experimental and simulated) as a function of time in vitro heart using acoustic power of 6 W. The spacing between the transducer and the heart tissue was 1 mm. The agreement between experimental and simulated results is excellent. Fig. 4B shows the lesion depth (experimental and simulated) as a function of power for a sonication duration of 35 s. The spacing between the transducer and heart tissue was also 1 mm. Based on the results of Fig. 4A it is concluded that in order to achieve 15 mm depth with acoustic power of 6 W exposure duration of 60 s is needed. From Fig. 4B it can be shown that to create a lesion of 15 mm 35 s are needed at a power of 9 W (Fig. 4B).

Fig. 5A shows a photograph of two lesions created in lamb heart in vitro at a plane perpendicular to the transducer beam axis. The power used was 6 W for 60 s. Note that the area of this lesion is slightly above the transducer area which is 10 × 3 mm². During

<table>
<thead>
<tr>
<th>Parameter</th>
<th>In vivo values</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Heart</td>
</tr>
<tr>
<td>Thermal conductivity (W m⁻¹°C⁻¹)</td>
<td>0.537*</td>
</tr>
<tr>
<td>Specific heat (J kg⁻¹°C⁻¹)</td>
<td>3720*</td>
</tr>
<tr>
<td>Initial temperature (°C)</td>
<td>37</td>
</tr>
<tr>
<td>Perfusion (kg m⁻³ s⁻¹)</td>
<td>1.42</td>
</tr>
<tr>
<td>Density (kg m⁻³)</td>
<td>1060</td>
</tr>
<tr>
<td>Attenuation (Np m⁻¹ MHz⁻¹)</td>
<td>4.1*</td>
</tr>
</tbody>
</table>

Gross anatomy the lesion depth was close to 15 mm. Fig. 5B shows the MRI image of the result of Fig. 5A using T1-W FSE for TR of 500 ms. Note that the lesion appears much brighter than the heart tissue, resulting to good image contrast. Fig. 5C shows the MRI image of the result of Fig. 5A using T2-W FSE with TE of 80 ms. In the T2-W image the lesion appears darker compared to the heart tissue.

T1-W FSE was investigated further by using different TR (200–1000 ms with increment step of 100 ms) and T2-W FSE was investigated further by using different TE (20, 40, 60, 100, and 120 ms).

Fig. 4. (A) Depth of lesion (experimental and simulated) as a function of time in vitro heart using acoustic power of 6 W. The spacing between the transducer and heart tissue was 1 mm. (B), Depth of the lesion (experimental and simulated) as a function of power for sonication duration of 35 s. The spacing between the transducer and heart tissue was also 1 mm.

Fig. 5. (A) Photograph in vitro of two lesions created in lamb heart in vitro at a plane perpendicular to the transducer beam axis. The power used was 6 W for 60 s. (B). MRI image of the result of figure A using T1-W FSE for TR of 500 ms. (C). MRI image of the result of figure A using T2-W FSE with TE of 80 ms.
ratio (CNR) of T2-W FSE plotted against TE in lamb heart in vitro. Note that the CNR with T2-W FSE is much lower than T1-W FSE.

Fig. 7A shows a photo of a lesion created in the heart of lamb tissue in vitro using 7 W at 35 s. The shape and size is in very good agreement with the lesion predicted by the simulation model (Fig. 7B). The experimental lesion depth was 10 mm, whereas the simulated lesion depth was 9.5 mm.

Fig. 8 shows a photo of a lesion created in the heart of rabbit in vivo using 6 W at 120 s. The area of the lesion in a plane parallel to the transducer follows the area of the transducer \(3 \times 10 \text{ mm}^2\). Despite of the motion of the heart and the high perfusion in the heart, it was feasible to obtain lesion size similar to what was obtained in the in vitro case \(3 \times 10 \text{ mm}^2\), however the penetration depth in the tissue was 12 mm compared to 15 mm (in vitro case). The thickness of this particular heart was 19 mm. The lesion was measured after the animal was sacrificed using a digital calliper.

Lastly, Fig. 9 shows the simulated depth vs. time at various levels of acoustic power. This figure essentially shows the main outcome of this study in one graph. It shows transducer (lesion depth) vs. the two main user-selected parameters (power and time). Note that the higher the power, the higher the rate of increase of lesion depth with time.

Fig. 10 shows the lesion depth created by the transducer using 6 W for 60 s for the various in vivo experiments. The mean lesion depth (excluding 2 experiments that failed due to possibly bad coupling) was 11.93 mm with a standard deviation of 0.62 mm.

4. Discussion

This paper presents a flat rectangular MR compatible transducer which can be used in the future to treat arrhythmias. This transducer was evaluated in lamb heart in vitro and in rabbit heart in vivo. A major part of the paper was to evaluate the performance of this transducer using a thermal simulation model. In this study, it is shown that the experimental and simulated results are in reasonable agreement, indicating that the theoretical model can be used to give guidelines for this type of transducer.

The target of this study was to create lesions of depth of at least 15 mm. This depth is sufficient to destroy unwanted electrical sig-
frequency transducer could be more beneficial [17]. Since this a transducer operating at the frequency of 5 MHz although a higher size in cardiac tissue is larger than the lesions created with other ablation techniques, such as radio frequency [13]. We have chosen shows that with this therapeutic ultrasound technology, the lesion however the ablation time (1 min per lesion) is low compared to necessarily so does the temperature. As a result, the lateral conduction if deeper in the tissue the intensity drops, and necessarily maximized because at low TRs, the difference between signal intensity of increased acoustic power, the exposure duration needed to achieve the same lesion dimensions deep in the tissue is decreased. In future clinical trials, this transducer can be attached to a catheter, and with MRI it will be easy to guide and monitor treatment. MRI has the added advantage that offers a better contrast between thermal necrosis and cardiac tissue than ultrasound or X-rays.

The size of the lesion in a plane parallel to the transducer face is shown in Fig. 7A (experimental) or Fig. 7B (simulated) is $3 \times 10 \text{ mm}^2$ (corresponds to the area of the transducer). At a deeper level, the size of the lesion becomes smaller. This is attributed to the fact that deeper in the tissue the intensity drops, and necessarily so does the temperature. As a result, the lateral conduction effect is lower and therefore the lesion size becomes smaller. If one needs a deeper lesion, then either the power or time has to be increased, taking care not to exceed temperatures above 100 °C. Increased ablation time means increased treatment time, however the ablation time (1 min per lesion) is low compared to the time needed to place in future clinical trials a catheter in the heart (30–60 min).

The transducer used produces lesions depth up to 15 mm. This shows that with this therapeutic ultrasound technology, the lesion size in cardiac tissue is larger than the lesions created with other ablation techniques, such as radio frequency [13]. We have chosen a transducer operating at the frequency of 5 MHz although a higher frequency transducer could be more beneficial [17]. Since this catheter is inserted in the body (i.e. it is consumable) going to higher frequency means a more expensive transducer. We use 5 MHz which is not very expensive and as we have proved the efficacy is not harmed.

There are many studies that predicted the size of the lesions as function of power and time (for example [54, 56, 57]). For example Damianou and Hynynen [56] evaluated the lesion size in dog muscle in vitro. Fagi et al. [57] performed a similar study in bovine in liver and Curiel et al. [54] have evaluated the lesion size in prostate. Therefore, for each application (in our case heart) and for every transducer (in our case planar) the lesion size has to be evaluated both using simulation and experimental models. This parametric information is very essential because when this technology is applied to humans the users should have an indicative knowledge of the lesion size.

The effect of perfusion is important especially using such long time duration (60 s). For example for 6 W and 60 s duration the in vitro experiments the depth of lesion is approximately 15 mm. Using the same exposure in vivo, the depth of lesion is 12 mm which is attributed to the perfusion effect [58].

In this paper the other major goal was to measure the CNR of FSE MRI sequences in detecting thermal lesions created by therapeutic ultrasound in heart. Both T1-W FSE and T2-W FSE have been proven successful for providing excellent contrast between rabbit heart and thermal lesion. The maximum contrast measured with T1-W FSE is approximately 22. With T1-W FSE the TR under which CNR is maximized is between 300 ms and 800 ms.

The trend of CNR vs. TR starts to increase then it becomes flat and then at high TRs it starts to decrease again. This trend is justified because at low TRs, the difference between signal intensity of lesion and tissue is low at short times and therefore CNR is lower. At higher TR the signal intensity of lesion and tissue reaches their maxima and therefore the signal difference is lower and hence the CNR drops again.

The maximum CNR measured with T2-W FSE is 13. With T2-W FSE the TE under which CNR is maximized is between 30 and 60 ms. The trend of CNR vs. TE starts to increase then it becomes flat and then at high TEs it starts to decrease again. The same explanation holds as in the case of T1-W FSE.

The cost of this prototype research unit (transducer, cabling and tubing for cooling) is approximately 1500 Euros. If the transducer, cabling and tubing for cooling are produced as a final product for clinical use in large quantities the cost could be close to 600 Euros. This figure puts the HIFU ablation catheter at a cost which is about 20% more expensive than the RF ablation catheter.

We still believe that with the use of MRI the treatment of arrhythmias has a clinical use (despite the complication mentioned...
by Metzner et al. 2010 [42]), because with the near real time imaging, the result of the therapy can be monitored. Therefore it is too early to abandon this technology for treating arrhythmias. Maybe in the future we may come to the conclusion that MRI cannot monitor the heating effects in the beating heart precisely, however this must be determined in future MRI studies.

4.1. Future work

In vitro MRI experiments do not provide much value due to the fact that the MRI response of a living tissue is different from ex vivo tissue responses. Thus, more in vivo work has to be done, especially regarding the MRI evaluation of thermal lesions. Additional MRI routines have to be explored because the FSE sequences used are long and sensitive to motion and could not be the optimum routines in a beating heart.

The key advantage of MRI is its ability to map temperature. However, this is difficult in a beating heart. In this paper there was no attempt to map the temperature in the heart tissue using MRI, and therefore this is a key future task.

References


