# Design and Analysis of a Whole Body Non-Contact Electromagnetic Stimulation Device with Field Modulation

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#### Abstract

This study describes a whole-body, non-contact electromagnetic stimulation device based on the concept of a conventional MRI Radio Frequency (RF) resonating coil, but at a much lower resonant frequency (100–150 kHz), with a field modulation option (0.5–100 Hz) and with an input power of up to 3 kW. Its unique features include a high electric field level within the biological tissue due to the resonance effect and a low power dissipation level, or a low Specific Absorption Rate (SAR), in the body itself. Because of its large resonator volume together with non-contact coupling, the subject may be located anywhere within the coil over a longer period at moderate and safe electric field levels. The electric field effect does not depend on body position within the resonator. However, field penetration is deep anywhere within the body, including the extremities where muscles, bones, and peripheral tissues are mostly affected. A potential clinical application of this device is treatment of chronic pain. Substantial attention is paid to device safety; this includes both AC power safety and exposure of human subjects to electromagnetic fields. In the former case, we employ inductive coupling which eliminates a direct current path from AC power to the coil. Our design enhances overall device safety at any power level, even when operated under higher-power conditions. Human exposure to electromagnetic fields within the coil is evaluated by performing modeling with two independent numerical methods and with an anatomically realistic multi-tissue human phantom. We show that SAR levels within the body correspond to International Electrotechnical Commission (IEC) safety standards when the input power level of the amplifier driver does not exceed 3 kW. We also show that electric field levels generally comply with International Commission on Non-Ionizing Radiation Protection safety standards if the input power level does not exceed 1.5 kW.

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## I. Introduction

In this present study, we have designed and constructed a whole-body non-contact, electromagnetics stimulation device based on the concept of an MRI Radio Frequency (RF) resonating coil. However, this has been accomplished at a much lower resonant frequency (100–150 kHz) and with a field modulation option (0.5–100 Hz) at input powers up to 3 kW. Due to a large resonator volume and its non-contact nature, the subject may be conveniently located anywhere within the resonating coil over a longer period of time at moderate and safe electric field levels. The electric field effect does not depend on a particular body position within the resonator. The field penetration is deep everywhere within the body including the extremities; muscles, bones, and peripheral tissues are mostly affected. Over a shorter period of time, the electric field levels may be increased to relatively large values with an amplitude of about 1 V/cm.

The primary potential clinical application for the device is treatment of chronic pain. Approximately 20% of the adult population report chronic pain and approximately one fifth of them have neuropathic pain characteristics [1–4]. Common forms of chronic pain include nociceptive pain, which involves stimulation of pain receptors secondary to tissue damage, or neuropathic pain, which involves injury or dysfunction of the central or peripheral nervous system [5]. Neuropathic pain is often rated as particularly intense and distressing and can have a significant negative impact on activities of daily living and quality of life [1–4]. A commonly prescribed treatment for chronic pain is opioids [6]. However, the use of opioids for pain relief is controversial owing to concerns about addiction and misuse [7].

One known yet not very efficient electromagnetic treatment of neuropathic and other pain conditions is Transcutaneous Electrical Nerve Stimulation, or TENS. It typically relies upon a portable unit which applies low-level electrical currents through electrodes attached to the skin. However, proper placement of electrodes and mitigating skin irritation are two major points of concern [4]. A well-known problem with electrodes attached to the skin is a strong regional current injection close to the contact area but not necessarily close to the deeper treatment region and may result in burning effects in the vicinity of electrodes where deep penetration is required. A number of systematic reviews of the effect of TENS on various painful conditions such as neuropathic pain, phantom limb pain, fibromyalgia, labor pain, rheumatoid arthritis, chronic lower back pain, etc. are available [1, 4, 8–12].

An alternative to electrical stimulation with contact electrodes is magnetic stimulation, which

induces electric field in the body via electromagnetic induction thus alleviating the issue of current shunting in the skin. The major goal of this paper is to introduce a distinctive, non-contact yet potentially powerful full-body electromagnetic stimulation device concept, including

- i. the underlying physical model in the form of an electromagnetic resonator;
- ii. the hardware design, preliminary measurements, and tests;
- iii. a detailed computational analysis of the electric fields induced in the body along with safety estimates.

The key idea of our stimulator is the creation of an initial rotating magnetic field that is routinely produced in resonant MRI RF coils, but for a completely different purpose, namely atomic spin excitation and RF signal acquisition. Furthermore, the resonant frequencies used in MRI RF coils are very high, typically  $64\,\mathrm{MHz}$  or higher, whereas our coil operates at much lower resonant frequencies ( $100-150\,\mathrm{kHz}$ ). Nonetheless, an RF birdcage coil [13, 14] seems to be an ideal starting point for our design. When the frequency becomes low as in the present case, the standard RF birdcage coil will possess very low inductance. Tuning such a birdcage toward resonance at low frequencies would require large capacitances. This, however, means a low Q-factor (quality factor Q is the "gain" of the resonator) and higher costs, as well as higher fabrication complexity [15, 16], and will restrict the use of the conventional birdcage coil to frequencies above at least a few megahertz. Different methods to overcome this difficulty have been suggested [15-20], but they are all limited to small-size coils.

Our design utilizes a unique large-scale low-pass birdcage coil architecture with a large number (144) of long rungs and bridging capacitors, and possesses a superior quality factor of approximately 300. The design does not use any magnetic materials; it is robust, lightweight, and portable. However, it achieves quite significant electric field levels within the body due to the resonance effect. This allows us to use relatively low input power levels and standard low-cost power electronic equipment. Although the resonant or carrier frequency is fixed, the electric field within the resonator volume can be amplitude modulated from approximately 0.5 Hz to 100 Hz.

The study is organized as follows. Section II describes a theoretical device model and specifies field distribution within the resonator. Section III describes hardware design, test, and functionality, including semiautomatic operation/tuning and representative continuous run times. Section IV provides computational results for the electric field distribution within the body obtained via two independent numerical methods. Section V discusses possible application scenarios as well as device modifications. Section VI concludes the paper.

### II. THEORETICAL MODEL OF THE STIMULATOR

# A. Concept of the magnetic stimulator. Two-dimensional analytical solution

Fig. 1 shows the anticipated device concept. An external uniform rotating (or circularly-polarized) magnetic flux density B with amplitude  $B_0$  is created in the transverse plane around a tissue volume, depicted in Fig. 1a. By Faraday's law of induction, this field excites a axial, rotating electric field E in free space or in a tissue volume, which is expressed in terms of the magnetic vector potential  $\mathbf{A}$ ,

$$\mathbf{E} = -\frac{\partial \mathbf{A}}{\partial t},\tag{1}$$

as shown in Fig. 1b; we set  $\mathbf{B} = \nabla \times \mathbf{A}$ . Thus, when a biological body is placed into this volume, a significant non-invasively excited electric field in the *axial* direction will appear parallel to the major peripheral nerves, spinal cord, long bones, major arteries, veins and other structures. This is in contrast to a solenoidal coil wound around the body creating electric fields and currents in the less desirable transverse plane.

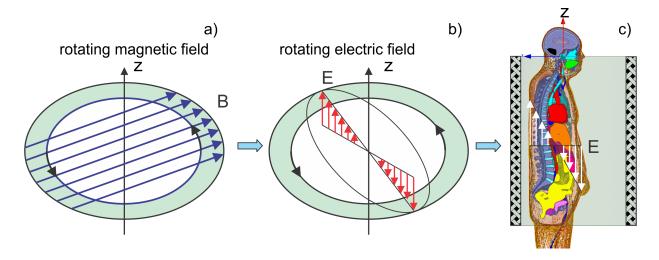


Fig. 1. Concept of non-invasive electric field excitation via the induction mechanism. a) A rotating magnetic field shown in excites the b) rotating electric field within the conductor and c) within the body.

The rotational character of the field also assures that not only one body cross-section (e.g. coronal or sagittal) will be subject to the electric field excitation, but the entire body volume.

In the ideal, two-dimensional case and for any conducting target with a strict cylindrical symmetry placed into the device, either homogeneous or not, the corresponding two-dimensional

problem, shown in Fig. 1a,b will have an exact analytical solution in the quasi-static (or eddy current) approximation. The electric field within the target is given by [21]

$$E_x = 0, \quad E_y = 0, \quad E_z = \omega B_0 r \cos(\omega t - \phi),$$
 (2)

$$\mathbf{J} = \sigma \mathbf{E} \tag{3}$$

where r is the radial distance from the coil axis in cylindrical coordinates,  $\omega$  is the angular frequency,  $\phi$  is an arbitrary phase, J is induced current density, and  $\sigma$  is the (local) medium conductivity, which is either constant or obeys cylindrical symmetry. Although Eqs. (2)–(3) might be used to roughly estimate the electric field in the body based on a cylindrically symmetric assumption, its actual value will deviate as shown below.

# B. Three-dimensional coil model

The external rotating magnetic flux density **B** is created using a volumetric resonator in the form of a low-pass birdcage coil. This design is well-known in the magnetic resonance radio-frequency coil community [13, 14]. We modified the design to operate at a much lower frequency of 100 kHz and with a very high loaded quality factor.

A computational model of a particular resonator constructed in this study is shown in Fig. 2a with the electric current distribution to scale. The coil has a diameter of 0.94 m and a length of 1.10 m; the coil resonates at 100 kHz or at 145 kHz depending on the values of the bridging capacitors. The coil consists of two rings (top and bottom) joined via multiple straight rungs, each bridged with a lumped capacitor at its center. The capacitors control the coil's resonant frequency. The resonating coil is fed via two lumped ports in quadrature, or using inductive coupling with two loops in quadrature as explained below.

From the modeling point of view, the resonant electric current in both rings at any fixed time instant behaves like a full period of a sine function of polar angle  $\varphi$ . This ring current distribution is shown in Fig. 2a. As time progresses, the ring current distribution shown in Fig. 2b rotates with angular frequency  $\omega$ . As a result, the time-domain ring current  $i(t,\varphi)$  in the top and bottom rings can be expressed in the form

$$i(t,\varphi) = \pm I_0 \cos(\omega t - \varphi + \pi/2), \quad \varphi \in [0,2\pi],$$
 (4)

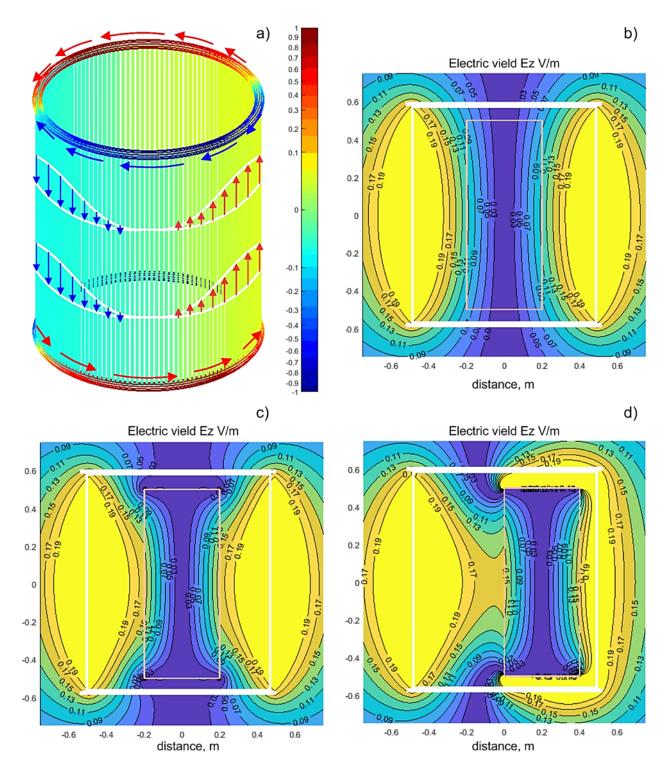


Fig. 2. Current distribution in the coil resonator and the associated electric field created by the coil when the ring current amplitude is 1 A for one resonant mode. a) Electric current distribution in the coil along with the current colorbar to scale; b) magnitude of the vertical electric  $E_z$  in V/m for an empty coil in the coronal plane; c) magnitude of the vertical electric  $E_z$  in V/m for the coil with a coaxial conducting cylinder  $0.4 \times 1$  m inside; c) magnitude of the vertical electric  $E_z$  in V/m for the coil with a conducting cylinder  $0.4 \times 1$  m shifted in the transverse plane inside.

where  $I_0$  is the current amplitude determined by the excitation power and by the quality factor (or the "gain") of the resonator.

The AC current in each rung shown in Fig. 2a does not change along its length. Simultaneously, at any fixed time instant, it also varies from rung to rung as a harmonic function of the polar angle  $\varphi$  with the full period corresponding to the ring circumference. This rung current distribution is shown in Fig. 2a. As time progresses, the rung current distribution shown in Fig. 2a also rotates with angular frequency  $\omega$ . Each individual time-domain rung current density  $j(t,\varphi)$  can be expressed in the form

$$j(t,\varphi) = \frac{I_0}{F}\cos(\varphi + \omega t),$$

$$F = \frac{1}{2} \sum_{m=0}^{\frac{N}{2} - 1} \sin\left(\frac{2\pi m}{N}\right), \quad \varphi \in [0, 2\pi]$$
(5)

where N is the total number of rungs. This form obeys the current conservation law, or Kirchhoff's Current Law (KCL), at every ring-rung junction.

The useful current, which creates a nearly constant horizontal rotating magnetic field  $B_r$  with amplitude  $B_0$  and axial rotating electric field  $E_z$  according to Eq. (2), is the rung current density  $j(t,\varphi)$ . Contributions of each rung add up in a constructive manner. The ring current, on the other hand, does not contribute to the axial (or vertical) electric field,  $E_z$ . However, it may create a strong transverse electric field very close to the rings.

It should be pointed out that Eqs. (4) and (5) describe the rotating current behavior, which is a combination of two elementary resonant modes. Each elementary mode does not rotate and appears as depicted in Fig. 2a. However, when excited in quadrature (with a 90 degree phase shift and a 90 degree excitation offset along the coil circumference), both modes combine to create the current distribution given by Eqs. (4) and (5) and the associated rotating electric field. The rotation phenomenon enable us to treat the entire body and not merely a singular component or region.

# C. Electric field model with and without a simple conducting object

Fig. 2b–d show the resulting electric field distribution in the coil (coronal plane) when the current amplitude  $I_0 = 1$  ampere in either ring given by Eq.(4). The results are given for one resonant mode is shown in Fig. 2a. Due to linearity, this result can simply be scaled for other excitation levels. Accurate field computations have been performed with the fast multipole

method described in [22]. The magnitude of the axial component  $E_z$  in V/m for an empty coil is shown in Fig. 2b. The electric field is indeed zero at the coil's center.

When a conducting object representing a load is inserted into the coil, the field distribution changes. Fig. 2c shows the distribution when a conducting cylinder with a diameter of 0.4 m and a length of 1 m is inserted into the coil along its axis. The particular conductivity value  $\sigma$  does not matter since only the conductivity contrast,  $(\sigma - \sigma_{air})/(\sigma + \sigma_{air})$ , is present in the solution [22]. This value is always unity since  $\sigma_{air} = 0$ .

An interesting and useful effect is observed in Fig. 2c: we see a "pulling" of the electric field into the cylinder close to the coil center. This is due to surface charges that appear at and near the cylinder tips. As a result, the electric field close to the cylinder surface at the center plane of the coil increases by nearly 36%.

Another remarkable observation (this effect is common in MRI RF coils) follows from Fig. 2d where the conducting cylinder has been shifted from the coil axis to the right by 0.2 m. While the electric field outside the conducting cylinder clearly changes, the field within the cylinder remains nearly the same, as observed in Fig. 2c. This may be explained as a result of the electric field being induced by the magnetic field, similar to eddy currents. Since the magnetic field is relatively homogeneous in the transverse plane of the coil, the induced electric fields in a load should not strongly depend on the transverse position of the load within the coil.

These rudimentary simulations allow us to establish two basic facts relevant for the analysis of realistic electric field distributions in a human body within the coil. First, we expect that the average transcutaneous electric field will be slightly higher than predicted by the air-filled coil model in dorsal, abdominal, and lumbar body regions. Second, we expect that the field within the body will not change significantly when the body is moved within the coil in the transverse plane; this circumstance seems to be quite useful from a practical point of view.

Obviously, an accurate field evaluation with a realistic human model is desirable; we will present our modeling results in Section IV.

## III. HARDWARE DESIGN AND TEST

# A. Power amplifier/driver

In order to create the rotating magnetic and electric fields as seen in Fig. 1a-b, two resonant modes are excited in the coil resonator. These modes display the same current distribution as

shown in Fig. 2a, but rotated by 90 degrees about the coil axis with respect to each other as well as having an additional temporal phase shift of 90 degrees.

To accomplish this, a custom designed class-D, high-efficiency, single-frequency power amplifier (PA), whose circuit schematic is presented in Fig. 3a, was constructed and prototyped as shown in Fig. 3b–c. The upper block in Fig. 3a is a class-S modulator, which is followed by two class-D output stages in quadrature exciting the two resonant modes. The PA has two outputs for each resonant mode and generates a harmonic power RF signal at a fixed carrier frequency at each output. This frequency is typically around 100 kHz. At the same time, the PA may be tuned to operate at any carrier frequency from 30 kHz to 300 kHz in the LF band. The PA operation, including variable power levels, an optional variable modulation frequency, and a semiautomatic patient-specific RF frequency tuning procedure, which is automated via a microcontroller board, can be seen in Fig. 3b.

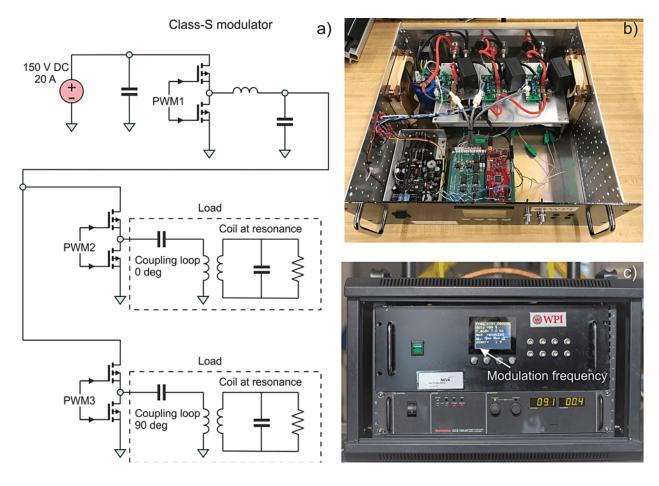


Fig. 3. a) High-level circuit schematic of the two-channel power amplifier; b) rackmount air-cooled assembly of the electronic hardware; c) amplifier display controlling output power and modulation frequency (if used).

The PA output stage is powered by a 3 kW Sorensen DCS 150-20 Variable Regulated DC (Direct Current) power supply seen at the bottom of Fig. 3c. When connected to a standard three-phase 208 VAC outlet, the max RF output power is about 2.9 kW, based on 3 kW DC power. Alternatively, when connected into a single-phase 240 VAC outlet, the max RF power reduces to about 2.3 kW based on 2.4 kW DC power.

Arbitrary *modulation* (pulse or CW) of the carrier signal with a maximum modulation frequency component of 1 kHz is available via the modulator. The modulation bandwidth is limited mainly by the coil envelope time constant of about 1 ms. Typical modulation is sinusoidal in the 0.5 Hz to 100 Hz range, generated by the PA firmware.

The PA also monitors its output power and load impedance. It uses this information to automatically adjust the carrier frequency in a narrow band such that the output power remains on target. The amplifier cost including the DC power supply is under \$10,000. The prototype 100 kHz PA was assembled in a rackmount case shown in Fig. 3b–c.

The reason for designing a custom, fixed-frequency PA is the lack of an affordable and appropriate commercial model. Industrial low frequency RF power supplies, e.g., Comdel's CLB3000, are costly and require a matched  $50\Omega$  load. Because our load impedance varies widely with frequency, keeping the load matched is a challenge. It would require load impedance monitoring and fine frequency control (potentially difficult with a commercial unit), and/or a software-controlled matching network (costly). Additionally, generating two outputs in quadrature would require either a  $90^{\circ}$  hybrid (another costly component), or phase-locking two commercial PAs at a  $90^{\circ}$  phase difference, which can be difficult. Finally, the majority of commercial PAs require water cooling, whereas our PA relies on air cooling. One disadvantage of our custom design is its unknown reliability, a factor that will be proven over time.

# B. Coupling and matching power amplifier to the resonating coil

The amplifier is coupled to the resonating coil inductively via two proximate loops. One such loop is shown in Fig. 4c. Apart from certain technical advantages of the inductive coupling, this methodology assures that there is no direct current path from the AC power outlet to the coil. This design enhances overall device safety at any power level, including high-power operation.

The matching network for a single coil port is shown in Fig. 4a. Two ports with identical matching networks are located 90° apart around the coil structure, as shown in Fig. 4b. The

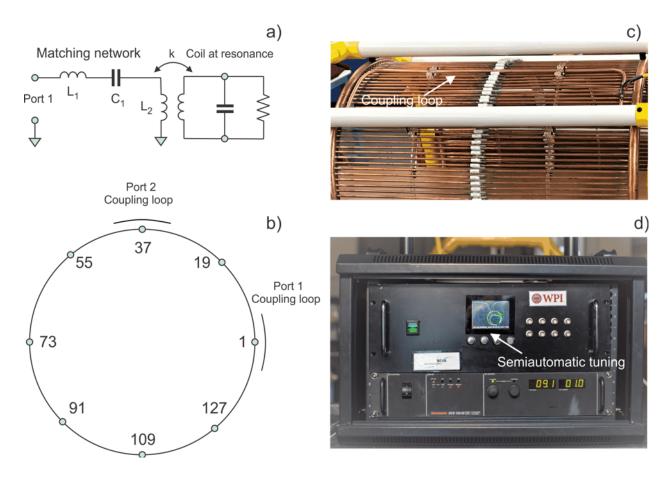


Fig. 4. a) Matching and tuning network of the power amplifier; b) assembly of two coupling-loop feed around the coil circumference with 144 rungs; c) non-contact inductive coupling of the power amplifier to the coil resonator at one port; d) Smith chart/reflection coefficient display of the power amplifier controller used for semiautomatic tuning at any desired time instant.

port matching network consists of a series capacitance  $C_1$ , series inductance  $L_1$ , and the fixed inductance  $L_2$  of the inductive loop shown in Fig. 4c.

The matching network is tuned such that the load looks mostly resistive over a small frequency band around the coil's resonance. For example, the load reactance stays quite low from 99.85 kHz to 100.15 kHz, while the resistance varies from  $1.3\Omega$  to  $6\Omega$ . Because the coil resonance shifts as the coil heats up, the operating frequency must be actively adjusted to compensate for this change, or the output power will vary.

Another important safety feature of the matching network is its benign behavior when subjected to a step response in output power. The matching network avoids large spikes in PA output current while energy is building up in the resonating coil.

Finally, the matching network presents a sufficiently inductive impedance to higher harmonics of the PA output voltage. This protects the output stage, and ensures that voltage transitions

occur when the output current is low, thereby improving efficiency. The efficiency of the PA with the expected load is estimated to be greater than 90% over a wide output power range.

# C. Tuning procedure

The semiautomatic tuning procedure ensures that the reflection coefficient of both modes stays below  $-25 \, \mathrm{dB}$  when matched to the maximum-power coil impedance of  $Z_0 = 2.5 \Omega$  and that both resonances are within a 20 Hz band. The primary adjustable components are the series capacitance  $C_1$  in Fig. 4a and the coil rung capacitors at the numbered locations in Fig. 4b.

The tuning procedure is controlled and guided by the Smith chart/reflection coefficient display of the PA controller seen in Fig. 4d. It includes a number of well-defined steps, and is applied to the coil at its designated operating location in an effort to account for the presence of large nearby metal objects. The tuning procedure is simple to perform.

# D. Coil assembly, complete device setup, and operation

A resonator coil prototype made of thin-walled, light copper tubing was constructed; it is shown in Fig. 5a–b. Tubing thickness was kept at 0.75 mm or greater to avoid excessive eddy current losses in the copper. The capacitor size in Fig. 5a is relatively large since those components must operate at significant currents levels, up to 15 A RMS per rung for a maximum power of 3 kW, and at large voltages, up to 240 V RMS for a maximum power of 3 kW across the capacitor. The total coil weight without the frame is approximately 120 lbs (54 kg).

This durable coil prototype was then framed, augmented with a horizontal bed, and placed horizontally to enable a subject to rest in the coil as shown in Fig. 5a, which simultaneously shows the complete device setup. The entire coil frame is portable. The distance between the PA, which is connected to the inductive coupling loops of the coil via two isolated cables, can vary from 1 to 3 meters, although larger distances may be possible. As mentioned above, there is no direct ohmic current path from the AC power outlet to the coil which is an important safety feature.

An operator sets the power level, the modulation frequency, and performs RF tuning at the beginning of the resonator operation and for a particular coil load. Furthermore, an automatic RF tuning procedure has been implemented that takes coil heating into account. At the maximum power level, the ring conductors of the coil heat up to approximately 70–75°C at continuous operation.

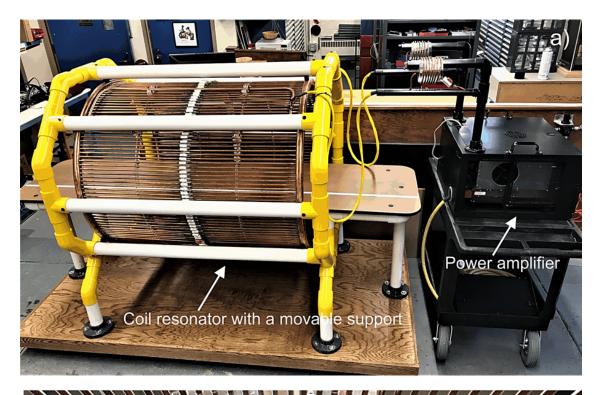




Fig. 5. a) Complete framed and movable device setup; b) coil resonator assembly.

Continuous coil operation at the maximum input power of 3kW was tested multiple times with an uninterrupted operation time of up to 2 hours and with a cumulative operating time in excess of 100 hours.

# E. Quality factor of the resonator and the magnetic field strength

The achievable field strength in the coil is determined by three factors: the strength of the PA, the quality factor Q or the "gain" of the resonator and the coil volume. When the quality factor is high, large field values within the coil can be achieved at a modest input power.

When measured across one of its rung capacitors, the birdcage coil behaves like a parallel resonator in a narrow frequency range around the resonant mode. Using a setup with a signal generator and oscilloscope, the resonator's quality factor has been estimated in the form:

$$Q = \frac{f_0}{(f_{\rm U} - f_{\rm L}) \left(1 - \frac{V_1}{V_0}\right)},\tag{6}$$

where  $f_0$  is the resonance frequency and voltage  $V_0$  is the open-circuited generator voltage. Voltage  $V_1$  is measured at resonance (where it is maximized),  $f_L$  and  $f_U$  are the lower and upper frequencies, respectively, where voltage  $V_1$  drops by 3 dB from its peak at resonance. This method is accurate in the high Q limit. The experimental data for 145 kHz is given in Table I. Table I reports a Q-factor value of about 300 at 145 kHz and a minimum difference between loaded (with a human body) and unloaded coil, which is to be expected at this low frequency. This value agrees with the theoretical/simulation prediction to within 10%. A similar quality factor (about 290) was obtained at 100 kHz.

TABLE I MEASURED Q-factors for the coil resonator at 145 kHz. Generator's voltage is 1 V RMS. The load is a 200 lb. subject.

Coil	$V_1$ (V RMS)	$f_0$ (kHz)	$f_{\rm L}$ (kHz)	f <sub>U</sub> (kHz)	$\overline{Q}$
Unloaded	0.6224	145.30	144.567	145.875	295.8
Loaded	0.6213	145.28	144.566	145.879	292.2

The established quality factor value is superior to the values reported in the literature for known low-frequency resonator coils (used for low-field MRI) in Table II. Note that all listed competitors have a much smaller coil size/volume and typically a lower quality factor.

TABLE II
CHARACTERISTICS OF EXISTING LOW-FREQUENCY RF COILS GIVEN FOR COMPARISON WITH THE PRESENT RESONATOR PROTOTYPE.

Ref. #	Coil type	Frequency (kHz)	Q (unloaded)
[15]	Wound birdcage coil (84 mm long and has a	386	180
	diameter of 73 mm)	300	100
[16]	Wound birdcage coils and a solenoid.	238/425	100–280
	The diameter and the length of the coils are 70 mm	230/423	100-200
[17]	27 turn saddle coil made of Litz wire with 8 cm diameter	373	105
[18]	4-Coil Whiting–Lee configuration, 33 cm long	83.6	100
[19]	Solenoidal coils; 6–46 cm in length and	210/275	60–30,
	4–52 cm in diameter	210/2/3	reduced Q
[20]	Cylindrical saddle-shaped loops (5 saddle pairs of	87	NA
	10 turns each), coil diameter is 26 mm	07	INA

It is important to point out again that the quality factor in Table I is weakly affected by body loading, in contrast to conventional high-frequency MRI RF coils. This observation, also mentioned in [16], is a critical advantage of the present electromagnetic stimulator. As to the RF power losses, they are mostly in the coil itself and not in the human body.

B-field measurements have been performed via a calibrated single-axis coil probe located at the coil axis. The measured and theoretical results differ by no more than 10%.

# IV. DEVICE SAFETY

### A. Method of analysis

Safety estimates rely upon the levels of the magnetic field, electric field, and the so-called Specific Absorption Rate (SAR) within the body. The SAR is the heat rate in the body rising due to an imposed electromagnetic field. At the low frequencies considered in this study, the magnetic field is weakly perturbed by the body presence and can therefore be measured in air. However, SAR and electric field measurements cannot be performed easily for human subjects *in vivo*. SAR and device performance estimates are typically derived and accepted today from Computational Electromagnetics (CEM) simulations performed with detailed virtual humans [23]. In this study, we use the multi-tissue CEM phantom VHP-Female v. 5.0 (female/60 year/162 cm/88 kg, obese) [23, 24], derived from the cryosection dataset archived within the Visible Human Project® of the US National Library of Medicine. The phantom includes about 250 individual tissues and is augmented with material property values from the

IT'IS database [25]. The average-body conductivity is assigned as 0.25 S/m, which reflects a mixture of muscle and fat.

The primary CEM software used in this study is the accurate commercial FEM solver AN-SYS® Electromagnetic Suite 18.2.0 with rigorous adaptive mesh refinement. In addition, and for verification purposes, we employ an in-house boundary element fast multipole method (BEM-FMM) described in [22]. In the latter case, a higher surface resolution can be achieved and the original surface phantom model can be refined and smoothed from approximately 0.5 M triangles to 3.5 M triangles. The human model is placed in the coil at the shoulder landmark, as shown in Fig. 6, so that the top of the shoulder coincides with the ring plane. Other configurations have also been considered. Results obtained with both software packages differ by no more than 2% in the unloaded coil and by no more than 25–50% in the coil loaded with the multi-tissue human body. The latter deviation may be explained by somewhat different surface meshes.

Below we report simulations at two power levels:  $1.5 \,\mathrm{kW}$  input power and  $3 \,\mathrm{kW}$  input power. The first power level is the half power level of the amplifier driver; the second power level corresponds to full power. At full power level, the amplitude of the resonant ring current  $I_0$  in Eqs. (4)–(5) reaches 603 A, while the amplitude of the rung current reaches 26 A.

# B. Electric field levels

Guidelines from the International Commission on Non-Ionizing Radiation Protection or IC-NIRP, see Table 2 of [26], require the occupational exposure to a time-varying electric field to be limited by a value of 27 V/m RMS at 100 kHz and by a value of 39 V/m RMS at 145 kHz (the so-called basic restrictions [26]). These restrictions are mainly due to the limits on peripheral nerve stimulation.

Fig. 7 shows the simulated RMS levels of the electric field in the body at 100 kHz and at the input power level of 1.5 kW obtained via the BEM-FMM simulations. We observe that, at half power level of the amplifier driver, the fields everywhere in the body do not generally exceed 30 V/m RMS and are thus within the ICNIRP guidelines. As expected, higher field levels are observed closer to the surface; the field gradually decreases toward the center of the body.

Quantitative estimates of the average electric field for every particular tissue obtained via ANSYS Electromagnetic Suite 18.2.0 are given in Table III. Note the lower electric fields in the intracranial volume. Additionally, we observe higher electric fields in the individual body

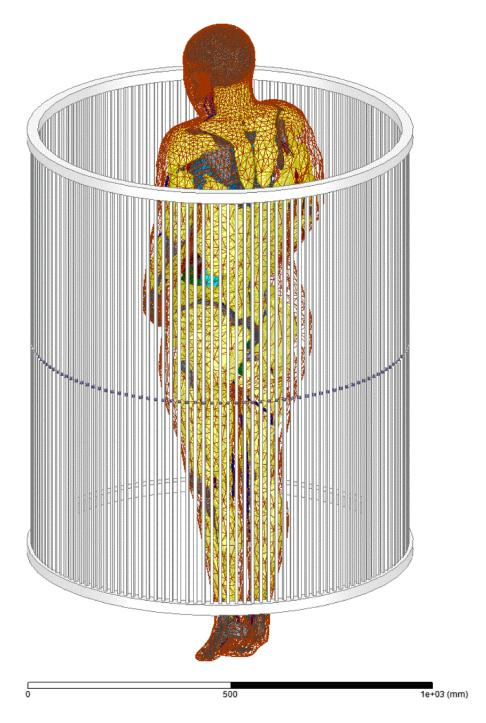


Fig. 6. Multi-tissue CEM phantom VHP-Female v. 5.0 within the resonant coil (ANSYS 18.2.0).

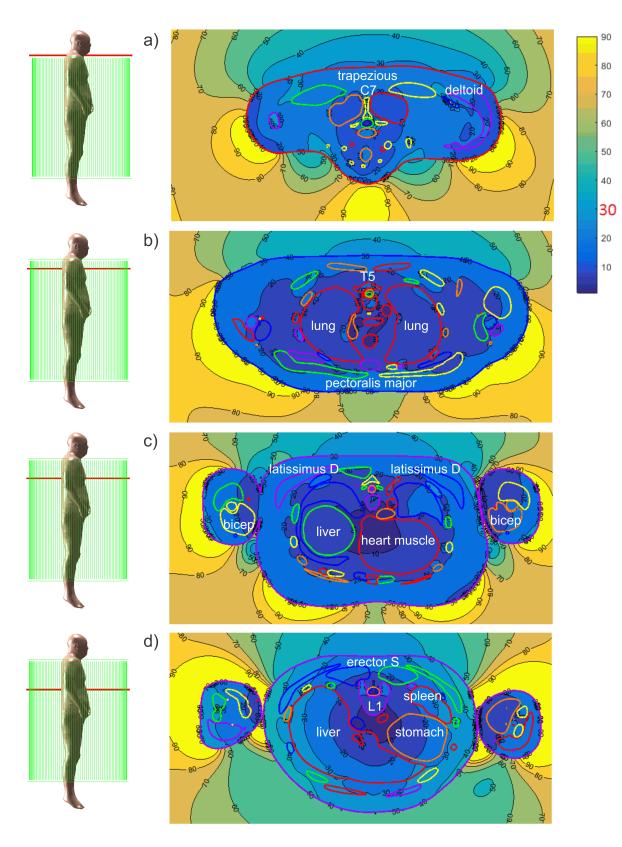


Fig. 7. Complex rms magnitude of the electric field (V/m) at 1,500 W input power.

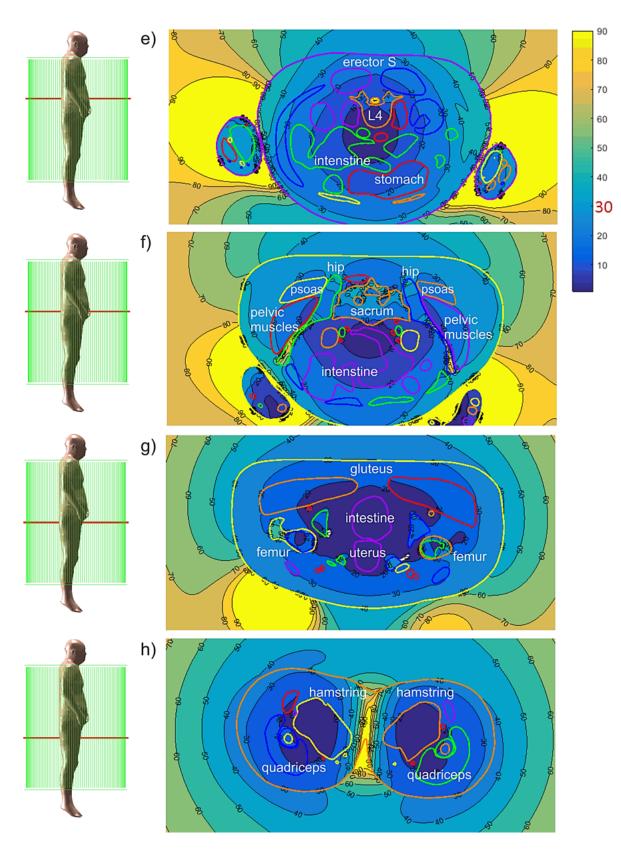


Fig. 7. (cont.). Complex rms magnitude of the electric field (V/m) at 1,500 W input power.

muscles. It is also interesting to observe that the fields in bone may be quite high, in particular in the femur and pelvic bones.

However, the computed local electric fields may considerably exceed the values reported in Table III, in particular by 1.5–6 times. These peak values are less accurate. One potential source of the numerical error is insufficient resolution of lengthy and time-consuming full-body computations very close to the interfaces where higher fields are usually observed.

TABLE III: Computed electric field levels (V/m RMS) in every individual tissue at 1.5 kW input power (ANSYS® Electromagnetic Suite 18.2.0).

Mesh	Tissue	Avg. E Field (V/m RMS)	Mesh	Tissue	Avg. E Field (V/m RMS)
1	Air Internal Maxillary Sinus Left	7.7	39	Cuneiform Medial right	0.6
2	Air Internal Maxillary Sinus Right	6.9	40	discC02C03	10.3
3	Arteries	10.5	41	discC03C04	11.6
4	Bladder	28.0	42	discC04C05	14.3
5	C01	14.5	43	discC05C06	16.9
6	C02	13.6	44	discC06C07	19.0
7	C03	14.8	45	discC07T01	20.6
8	C04	18.5	46	discL01L02	13.9
9	C05	21.7	47	discL02L03	11.5
10	C06	26.1	48	discL03L04	8.4
11	C07	29.6	49	discL04L05	4.7
12	Calcaneous left	0.6	50	discL05L06	7.1
13	Calcaneous right	1.1	51	discL06S00	13.8
14	Cartilage1 Left	18.5	52	discT01T02	20.2
15	Cartilage1 Right	19.9	53	discT02T03	17.6
16	Cartilage2 Left	19.7	54	discT03T04	17.9
17	Cartilage2 Right	20.3	55	discT04T05	17.4
18	Cartilage3 Left	21.2	56	discT05T06	15.8
19	Cartilage3 Right	20.9	57	discT06T07	15.2
20	Cartilage4 Left	22.5	58	discT07T08	14.4
21	Cartilage4 Right	21.8	59	discT08T09	13.6
22	Cartilage5 Left	24.3	60	discT09T10	13.2
23	Cartilage5 Right	22.6	61	discT10T11	12.5
24	Cartilage6 Left	36.3	62	discT11T12	13.2
25	Cartilage6 Right	35.9	63	discT12L01	13.4
26	Cerebellum	1.4	64	Eye Left	5.0
27	Clavicle left	55.6	65	Eye Right	5.3
28	Clavicle right	34.9	66	Feet1Phalange left	0.5
29	Coccyx	42.6	67	Feet1Phalange right	0.4
30	CSF OuterShell	3.5	68	Feet2Phalange left	0.4
31	CSF Ventricles	0.4	69	Feet2Phalange right	0.4
32	Cuboid Left	0.9	70	Feet3Phalange left	0.3
33	Cuboid Right	0.6	71	Feet3Phalange right	0.4

34	Cuneiform Intermediate left	1.3	72	Feet4Phalange left	0.4
35	Cuneiform Intermediate right	0.5	73	Feet4Phalange right	0.6
36	Cuneiform Lateral left	1.1	74	Feet5Phalange left	0.4
37	Cuneiform Lateral right	0.4	75	Feet5Phalange right	0.7
38	Cuneiform Medial left	1.3	76	Femur Bone Marrow Left	7.1
77	Femur Bone Marrow Right	8.6	117	Humerus right	23.1
78	Femur left	69.3	118	Intestine	20.6
79	Femur right	83.7	119	Jaw lower	10.0
80	Fibula left	5.9	120	Kidney left	29.3
81	Fibula right	5.4	121	Kidney right	27.2
82	Grey Matter Spinal Cord	1.5	122	L01	27.9
83	Hands1 1Phalange left	10.2	123	L02	25.3
84	Hands1 1Phalange right	9.5	124	L03	22.1
85	Hands1 2Phalange left	9.5	125	L04	19.0
86	Hands1 2Phalange right	12.1	126	L05	14.4
87	Hands1 3Phalange left	12.2	127	L06	17.9
88	Hands1 3Phalange right	13.4	128	Liver	29.8
89	Hands2 1Phalange left	7.6	129	Lungs	19.4
90	Hands2 1Phalange right	7.1	130	Median Nerve left	11.6
91	Hands2 2Phalange left	8.1	131	Median Nerve right	13.0
92	Hands2 2Phalange right	8.5	132	Muscle Bicep left	11.9
93	Hands2 3Phalange left	7.1	133	Muscle Bicep right	12.9
94	Hands2 3Phalange right	9.5	134	Muscle Calf left	5.0
95	Hands3 1Phalange left	6.4	135	Muscle Calf right	5.2
96	Hands3 1Phalange right	6.2	136	Muscle Deltoid left	18.7
97	Hands3 2Phalange left	8.1	137	Muscle Deltoid right	19.3
98	Hands3 2Phalange right	8.6	138	Muscle Erector spinae left	26.8
99	Hands3 3Phalange left	9.3	139	Muscle Erector spinae right	26.9
100	Hands3 3Phalange right	11.0	140	Muscle Forearm Extensors left	6.9
101	Hands4 1Phalange left	7.2	141	Muscle Forearm Extensors right	8.6
102	Hands4 1Phalange right	6.9	142	Muscle Forearm Flexors left	6.9
103	Hands4 2Phalange left	10.4	143	Muscle Forearm Flexors right	7.2
104	Hands4 2Phalange right	10.0	144	Muscle Gluteus left	27.9
105	Hands4 3Phalange left	11.1	145	Muscle Gluteus right	27.2
106	Hands4 3Phalange right	10.7	146	Muscle Hamstring left	18.9
107	Hands5 1Phalange left	9.0	147	Muscle Hamstring right	19.1
108	Hands5 1Phalange right	10.3	148	Muscle Latissimus Dorsi left	36.6
109	Hands5 2Phalange left	11.0	149	Muscle Latissimus Dorsi right	38.5
110	Hands5 2Phalange right	12.6	150	Muscle Neck Combined left	13.6
111	Hands5 3Phalange left	10.3	151	Muscle Neck Combined right	13.4
112	Hands5 3Phalange right	12.1	152	Muscle Obliques left	39.5
113	Heart Muscle	14.2	153	Muscle Obliques right	40.1
114	Hip left	60.0	154	Muscle Pectoralis major left	21.7
115	Hip right	61.4	155	Muscle Pectoralis major right	20.9
116	Humerus left	20.7	156	Muscle Pectoralis minor left	19.2
157	Muscle Pectoralis minor right	18.6	194	Ribs left8	47.3
158	Muscle Pelvic Combined left	25.7	195	Ribs left9	46.1
159	Muscle Pelvic Combined right	25.0	196	Ribs left10	48.5

160	Muscle Psoas left	13.9	197	Ribs left11	51.7
161	Muscle Psoas right	13.9	198	Ribs left12	39.0
162	Muscle Quadriceps left	20.2	199	Ribs right1	29.6
163	Muscle Quadriceps right	19.9	200	Ribs right2	26.2
164	Muscle Rectus Abdominis left bottom	32.3	201	Ribs right3	25.2
165	Muscle Rectus Abdominis left middle	34.9	202	Ribs right4	26.1
166	Muscle Rectus Abdominis left top	39.1	203	Ribs right5	27.2
167	Muscle Rectus Abdominis right bottom	32.5	204	Ribs right6	29.9
168	Muscle Rectus Abdominis right middle	35.4	205	Ribs right7	35.6
169	Muscle Rectus Abdominis right top	38.1	206	Ribs right8	43.2
170	Muscle Sartorius left	18.6	207	Ribs right9	53.9
171	Muscle Sartorius right	17.5	208	Ribs right10	58.9
172	Muscle Tibialis Anterior left	6.2	209	Ribs right11	56.9
173	Muscle Tibialis Anterior right	5.8	210	Ribs right12	40.9
174	Muscle Trapezius left	23.6	211	Sacrum	45.7
175	Muscle Trapezius right	24.0	212	Scapula left	38.2
176	Muscle Tricep left	12.0	213	Scapula right	38.8
177	Muscle Tricep right	14.0	214	Skin Shell	27.8
178	Navicular left	1.8	215	Skull	22.8
179	Navicular right	0.7	216	Sphenoid	8.9
180	Patella left	24.3	217	Spleen	33.9
181	Patella right	22.6	218	Sternum	25.2
182	Peripheral Nerve left	17.1	219	Stomach	22.6
183	Peripheral Nerve Right	14.1	220	T01	28.4
184	Pubic Symphysis	32.1	221	T02	27.2
185	Radial Nerve left	14.6	222	T03	27.2
186	Radial Nerve right	12.4	223	T04	26.9
187	Ribs left1	26.4	224	T05	25.7
188	Ribs left2	30.1	225	T06	25.5
189	Ribs left3	26.4	226	T07	26.2
190	Ribs left4	26.7	227	T08	26.4
191	Ribs left5	28.3	228	T09	26.9
192	Ribs left6	31.5	229	T10	26.1
193	Ribs left7	37.3	230	T11	26.4
231	T12	27.5	240	Trabecular upper right	0.9
232	Talus left	1.3	241	Trachea Sinus	12.4
233	Talus right	0.6	242	Ulna Radius left	8.1
234	Tibia left	8.3	243	Ulna Radius right	7.8
235	Tibia right	7.9	244	Uterus	17.3
236	Tongue	5.2	245	Veins lower	12.5
237	Trabecular lower left	0.5	246	Veins upper	12.4
238	Trabecular lower right	0.8	247	White Matter	1.0
239	Trabecular upper left	0.7			•

# C. SAR levels

The body-averaged or the whole-body (global-body)  $SAR_{body}$  is given by averaging the local SAR over the entire body volume. In terms of the complex field phasor  $E(\mathbf{r})$ , one has

$$SAR_{body} = \frac{1}{V_{body}} \int_{V_{body}} \frac{\sigma(\mathbf{r})}{2\rho(\mathbf{r})} \sqrt{\mathbf{E}(\mathbf{r}) \cdot \mathbf{E}(\mathbf{r})^*} \, dV$$
 (7)

Here,  $\sigma(\mathbf{r})$  is the local tissue conductivity and  $\rho(\mathbf{r})$  is the local mass density. At full power of 3 kW and positioned at the shoulder landmark, the global-body SAR computed via ANSYS Electromagnetic Suite 18.2.0 is 0.25 W/kg. Thus, the total power dissipation in the body does not exceed 30 W, i.e., 1% of the total power. The same percentage ratio is valid at half input power.

The second critical estimate is  $SAR_{1g}$ , which is given by averaging over a contiguous volume with the weight of 1g,

$$SAR_{1g} = \frac{1}{V_{1g}} \int_{V_{1g}} \frac{\sigma(\mathbf{r})}{2\rho(\mathbf{r})} \sqrt{\mathbf{E}(\mathbf{r}) \cdot \mathbf{E}(\mathbf{r})^*} \, dV$$
 (8)

The maximum value of  $SAR_{1g}$ , in the body computed via ANSYS Electromagnetic Suite 18.2.0 at the full power of 3 kW and located at the shoulder landmark is 4.55 W/kg.

Although this last value might appear to be relatively high, it is still within the corresponding SAR limits in MRI machines [27, 28]. In particular, the major applicable MRI safety standard [28] issued by the International Electrotechnical Commission (IEC) and also accepted by the U.S. Food and Drug Administration in the normal mode (mode of operation that causes no physiological stress to patients) limits global-body SAR to 2 W/kg, global-head SAR to 3.2 W/kg, local head and torso SAR to 10 W/kg, and local extremity SAR to 20 W/kg [27]. The global SAR limits are intended to ensure a body core temperature of 39°C or less [27, 28].

## V. DISCUSSION

We envision several potential application scenarios of the present electromagnetic stimulation device, though others may certainly be feasible. First, it could be tested for chronic pain treatment as suggested in the introduction. For example, one potential target is the spinal cord. Epidural spinal cord stimulation (SCS) has been used for pain management in the past several decades [29]. However, SCS involves surgical implantation of a pulse generator device that delivers weak currents to nerve fibers of the spinal cord. A noninvasive alternative is transcutaneous spinal

direct current stimulation (tcDCS) [30], which delivers an average electric field of 0.15 V/m in the spinal cord between the electrodes. In comparison, our coil can induce an electric field in the spinal cord grey matter of approximately an order of magnitude higher compared to direct current stimulation.

Another potential application for our coil device is treatment of psychiatric disorders such as major depressive disorder. Repetitive transcranial magnetic stimulation (rTMS) was first shown to be efficacious for the treatment of depression in the mid-1990s [31] and subsequently cleared by the U.S. FDA in 2008 for treatment-resistant depression (TRD). Clinical rTMS uses a focal figure-of-8 coil to deliver electric field pulses at the dorsal lateral prefrontal cortex at an intensity sufficient to induce action potentials in the underlying neurons. Several strategies using nonfocal (whole brain) subthreshold stimulation are currently being explored for the treatment of TRD. For example, it was reported that low-field (< 1 V/m) stimulation delivered using a MRI-gradient type coil had a rapid mood-elevating effect in animals and bipolar patients [32–35], and has been shown to affect brain glucose metabolism [36]. In addition, a system has been proposed, using a series of rotating permanent magnets to induce a small current in the brain in order to entrain neural oscillations, enhance cortical plasticity, normalize cerebral blood flow, and altogether ameliorate depressive symptoms [37]. An embodiment of our coil can be made smaller for efficient transcranial brain stimulation.

# VI. CONCLUSIONS

In this technical study we described a whole-body non-contact electromagnetic stimulation device based on the concept of a familiar MRI RF resonating coil, but at a much lower resonant frequency (100–150 kHz), with a field modulation option (0.5–100 Hz) and with an input power level of up to 3 kW. Its unique features include a high electric field level within the subject's biological tissue due to the resonant effect but at a low power dissipation, or SAR level, in the body itself.

Due to a large resonator volume and its non-contact nature, the subject may be conveniently located anywhere within the resonating coil over a prolonged period of time at moderate and safe electric field levels. The electric field effect does not depend on a particular body position within the resonator. The field penetration is deep everywhere in the body including the extremities; muscles, bones, and peripheral tissues are mostly affected. Over a shorter period of time, the

electric field levels could be increased to relatively large values with an amplitude of about 1 V/cm.

We envision treatment of chronic pain, and particularly neuropathic pain, as the primary potential clinical application for the device. The device enables whole-body coverage, which could be useful in the treatment of widespread pain conditions, such as painful polyneuropathy or fibromyalgia. In addition, a deeper tissue penetration can be achieved without causing side-effects caused by high current density in the skin associated with the traditional contact electrodes. Another potential application might include for example facilitation of chronic skin wound healing [38]. Medium to high electric field levels approaching or even exceeding 30 V/m RMS could likely be applied. It should be noted that these potential clinical applications are speculative and warrant empirical testing in the future.

Considerable attention has been paid to device safety including both the AC power safety and human exposure to electromagnetic fields. In the former case, we have used inductive coupling, which assures that there is no direct current path from the AC power outlet to the coil. This design enhances overall device safety at any power level, including high-power operation. As with more traditional MRI devices, no large metal objects should be located in the immediate vicinity of the coil.

Human exposure to the electromagnetic field within the coil has been evaluated by performing extensive modeling with two independent numerical methods and with an anatomically realistic multi-tissue human phantom. We have shown that the SAR levels within the body correspond to the safety standards of the International Electrotechnical Commission when the input power level of the amplifier driver does not exceed 3 kW. We have also shown that the electric field levels generally comply with the safety standards of the International Commission on Non-Ionizing Radiation Protection when the input power level of the amplifier driver does not exceed 1.5 kW.

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### REFERENCES

- [1] A. M. McDermott, T. R. Toelle, D. J. Rowbotham, C. P. Schaefer, and E. M. Dukes, "The burden of neuropathic pain: results from a cross-sectional survey," *Eur J Pain*, vol. 10, no. 2, pp. 127–135, 2006.
- [2] J. Kennedy, J. M. Roll, T. Schraudner, S. Murphy, and S. McPherson, "Prevalence of persistent pain in the U.S. adult population: new data from the 2010 national health interview survey," *J Pain*, vol. 15, no. 10, pp. 979–984, 2014.
- [3] M. I. Johnson, L. S. Claydon, G. P. Herbison, C. A. Paley, and G. Jones, "Transcutaneous electrical nerve stimulation (TENS) for fibromyalgia in adults," *Cochrane Database Syst Rev*, vol. 9, no. CD012172, 2017.
- [4] W. Gibson, B. M. Wand, and N. E. O'Connell, "Transcutaneous electrical nerve stimulation (TENS) for neuropathic pain in adults," *Cochrane Database Syst Rev*, vol. 9, no. CD11976, 2017.
- [5] B. Nicholson, "Differential diagnosis: nociceptive and neuropathic pain," *Am J Manag Care*, vol. 12, no. 9 Suppl, pp. S256–S262, 2006.
- [6] M. C. Rowbotham, L. Twilling, P. S. Davies, L. Reisner, K. Taylor, and D. Mohr, "Oral opioid therapy for chronic peripheral and central neuropathic pain," N Engl J Med, vol. 348, no. 13, pp. 1223–1232, 2003.
- [7] National Institute on Drug Abuse. Opioid overdose crisis. Access 2018-09-10. [Online]. Available: https://www.drugabuse.gov/drugs-abuse/opioids/opioid-overdose-crisis
- [8] I. Koca, A. Boyaci, A. Tutoglu, M. Ucar, and O. Kocaturk, "Assessment of the effectiveness of interferential current therapy and TENS in the management of carpal tunnel syndrome: a randomized controlled study," *Rheumatol Int*, vol. 34, no. 12, pp. 1639–1645, 2014.
- [9] L. Brosseau, M. G. Judd, S. Marchand, V. A. Robinson, P. Tugwell, G. Wells, and K. Yonge, "Transcutaneous electrical nerve stimulation (TENS) for the treatment of rheumatoid arthritis in the hand," *Cochrane Database Syst Rev*, vol. 3, no. CD004287, 2003.
- [10] A. Khadilkar, D. O. Odebiyi, L. Brosseau, and G. A. Wells, "Transcutaneous electrical nerve stimulation (TENS) versus placebo for chronic low-back pain," *Cochrane Database Syst Rev*, vol. 4, no. CD003008, 2008.
- [11] T. Dowswell, C. Bedwell, T. Lavender, and J. P. Neilson, "Transcutaneous electrical nerve stimulation (TENS) for pain management in labour," *Cochrane Database Syst Rev*, vol. 2,

- no. CD007214, 2009.
- [12] M. I. Johnson, M. R. Mulvey, and A. M. Bagnall, "Transcutaneous electrical nerve stimulation (TENS) for phantom pain and stump pain following amputation in adults," *Cochrane Database Syst Rev*, vol. 8, no. CD007264, 2015.
- [13] C. E. Hayes, W. A. Edelstein, J. F. Schenck, O. M. Mueller, and M. Eash, "An efficient, highly homogeneous radiofrequency coil for whole-body nmr imaging at 1.5 T," *J Mag Reson*, vol. 63, no. 3, pp. 622–628, 1985.
- [14] C. E. Hayes, "The development of the birdcage resonator: a historical perspective," *NMR Biomed*, vol. 22, no. 9, pp. 908–918, 2009.
- [15] H. M. Borsboom, T. Claasen-Vujci'c, H. J. Gaykema, and T. Mehlkopf, "Low-frequency quadrature mode birdcage resonator," *MAGMA*, vol. 5, no. 1, pp. 33–37, 1997.
- [16] T. Claasen-Vujcić, H. M. Borsboom, H. J. Gaykema, and T. Mehlkopf, "Transverse low-field RF coils in MRI," *Magn Reson Med*, vol. 36, no. 1, pp. 111–116, 1996.
- [17] A. Galante, R. Sinibaldi, A. Conti, C. De Luca, N. Catallo, P. Sebastiani, V. Pizzella, G. L. Romani, A. Sotgiu, and S. Della Penna, "Fast room temperature very low field-magnetic resonance imaging system compatible with magnetoencephalography environment," *PLoS One*, vol. 10, no. 12, e0142701, 2015.
- [18] I. Savukov, T. Karaulanov, A. Castro, P. Volegov, A. Matlashov, A. Urbatis, J. Gomez, and M. Espy, "Non-cryogenic anatomical imaging in ultra-low field regime: hand MRI demonstration," *J Magn Reson*, vol. 211, no. 2, pp. 101–108, 2011.
- [19] L. L. Tsai, R. W. Mair, M. S. Rosen, S. Patz, and R. L. Walsworth, "An open-access, very-low-field MRI system for posture-dependent 3He human lung imaging," *J Magn Reson*, vol. 193, no. 2, pp. 274–285, 2008.
- [20] V. V. Kuzmin, C. P. Bidinosti, M. E. Hayden, and P. J. Nacher, "An improved shielded RF transmit coil for low-frequency NMR and MRI," *J Magn Reson*, vol. 256, pp. 70–76, 2015.
- [21] S. N. Makarov, G. M. Noetscher, and A. Nazarian, *Low-Frequency Electromagnetic Modeling for Electrical and Biological Systems Using MATLAB*. New York: Wiley, 2015.
- [22] S. N. Makarov, G. M. Noetscher, T. Raij, and A. Nummenmaa, "A quasi-static boundary element approach with fast multipole acceleration for high-resolution bioelectromagnetic models," *IEEE Trans Biomed Eng*, 2018, Early Access: http://ieeexplore.ieee.org/document/8307445/.
- [23] S. N. Makarov, G. M. Noetscher, J. Yanamadala, M. W. Piazza, S. Louie, A. Prokop,

- A. Nazarian, and A. Nummenmaa, "Virtual human models for electromagnetic studies and their applications," *IEEE Rev Biomed Eng*, vol. 10, pp. 95–121, 2017.
- [24] H. Tankaria, X. J. Jackson, R. Borwankar, G. N. Srichandhru, A. Le Tran, J. Yanamadala, G. M. Noetscher, A. Nazarian, S. Louie, and S. N. Makarov, "VHP-Female full-body human CAD model for cross-platform FEM simulations: recent development and validations," in *Conf Proc IEEE Eng Med Biol Soc*, 2016, pp. 2232–2235.
- [25] P. A. Hasgall, E. Neufeld, M. C. Gosselin, A. Klingenböck, N. Kuster, N. Kuster, P. Hasgall, and M. Gosselin. (2015) IT'IS Database for thermal and electromagnetic parameters of biological tissues. [Online]. Available: www.itis.ethz.ch/database
- [26] International Commission on Non-Ionizing Radiation Protection, "Guidelines for limiting exposure to time-varying electric and magnetic fields (1 Hz to 100 kHz)," *Health Phys*, vol. 99, no. 6, pp. 818–836, 2010.
- [27] H. Homann, SAR prediction and SAR management for parallel transmit MRI. KIT Scientific Publishing, 2012.
- [28] Medical equipment–IEC 60601-2-33: Particular requirements for the safety of magnetic resonance equipment, International Electrotechnical Commission, International standard Std., 2010, Amendment 1 (2013), Amendment 2 (2015). [Online]. Available: https://webstore.iec.ch/publication/22705
- [29] K. Kumar, C. Toth, R. K. Nath, and P. Laing, "Epidural spinal cord stimulation for treatment of chronic pain—some predictors of success. a 15-year experience," *Surg Neurol*, vol. 50, no. 2, pp. 110–120, 1998.
- [30] S. R. Fernandes, R. Salvador, C. Wenger, M. de Carvalho, and P. C. Miranda, "Transcutaneous spinal direct current stimulation of the lumbar and sacral spinal cord: a modelling study," *J Neural Eng*, vol. 15, no. 3, p. 036008, 2018.
- [31] M. S. George, E. M. Wassermann, W. A. Williams, A. Callahan, T. A. Ketter, P. Basser, M. Hallett, and R. M. Post, "Daily repetitive transcranial magnetic stimulation (rTMS) improves mood in depression," *Neuroreport*, vol. 6, no. 14, pp. 1853–1856, 1995.
- [32] M. Fava, M. P. Freeman, M. Flynn, B. B. Hoeppner, R. Shelton, D. V. Iosifescu, J. W. Murrough, D. Mischoulon, C. Cusin, M. Rapaport, B. W. Dunlop, M. H. Trivedi, M. Jha, G. Sanacora, G. Hermes, and G. I. Papakostas, "Double-blind, proof-of-concept (POC) trial of Low-Field Magnetic Stimulation (LFMS) augmentation of antidepressant therapy in treatment-resistant depression (TRD)," *Brain Stimul*, vol. 11, no. 1, pp. 75–84, 2018.

- [33] M. L. Rohan, R. T. Yamamoto, C. T. Ravichandran, K. R. Cayetano, O. G. Morales, D. P. Olson, G. Vitaliano, S. M. Paul, and B. M. Cohen, "Rapid mood-elevating effects of low field magnetic stimulation in depression." *Biol Psychiatry*, vol. 76, no. 3, pp. 186–193, 2014.
- [34] M. Rohan, A. Parow, A. L. Stoll, C. Demopulos, S. Friedman, S. Dager, J. Hennen, B. M. Cohen, and P. F. Renshaw, "Low-field magnetic stimulation in bipolar depression using an MRI-based stimulator," *Am J Psychiatry*, vol. 161, no. 1, pp. 93–98, 2004.
- [35] W. A. Carlezon, Jr., M. L. Rohan, S. D. Mague, E. G. Meloni, A. Parsegian, K. Cayetano, H. C. Tomasiewicz, E. D. Rouse, B. M. Cohen, and P. F. Renshaw, "Antidepressant-like effects of cranial stimulation within a low-energy magnetic field in rats," *Biol Psychiatry*, vol. 57, no. 6, pp. 571–576, 2005.
- [36] N. D. Volkow, D. Tomasi, G. J. Wang, J. S. Fowler, F. Telang, R. Wang, D. Alexoff, J. Logan, C. Wong, K. Pradhan, E. C. Caparelli, Y. Ma, and M. Jayne, "Effects of low-field magnetic stimulation on brain glucose metabolism," *Neuroimage*, vol. 51, no. 2, pp. 623–628, 2010.
- [37] A. F. Leuchter, I. A. Cook, D. Feifel, J. W. Goethe, M. Husain, L. L. Carpenter, M. E. Thase, A. D. Krystal, N. S. Phillip, M. T. Bhati, W. J. Burke, R. H. Howland, Y. I. Sheline, S. T. Aaronson, D. V. Iosifescu, J. P. O'Reardon, W. S. Gilmer, R. Jain, K. S. Burgoyne, B. Phillips, P. J. Manberg, J. Massaro, A. M. Hunter, S. H. Lisanby, and M. S. George, "Efficacy and safety of low-field synchronized transcranial magnetic stimulation (sTMS) for treatment of major depression," *Brain Stimul*, vol. 8, no. 4, pp. 787–794, 2015.
- [38] A. Sebastian, S. A. Iqbal, J. Colthurst, S. W. Volk, and A. Bayat, "Electrical stimulation enhances epidermal proliferation in human cutaneous wounds by modulating p53-SIVA1 interaction," *J Invest Dermatol*, vol. 135, no. 4, pp. 1166–1174, 2015.