

Estimating the Electrical Conductivity of Human Tissue in Radiofrequency Hyperthermia Therapy

Estimación de la conductividad eléctrica del tejido humano en la terapia de hipertermia por radiofrecuencia

Jorge I. López-Pérez¹, and Leonardo A. Bermeo Varón²

ABSTRACT

The use of mathematical models to study complex systems such as physical and biological phenomena allows understanding their behavior, specifically regarding variables and parameters that are difficult to obtain. Additionally, studying optimization techniques has made it possible to approximate the characteristics of these systems by correlating numerical simulations and experimentation. Radiofrequency hyperthermia therapy for cancer treatment is currently under consideration for future medical applications. However, some of its properties are complex to measure, which could prevent their control. This is the case of electrical conductivity, which depends on the induction frequency and the tissue characteristics. In this paper, radiofrequency hyperthermia therapy was simulated via the finite element method. Then, an estimation of the electrical conductivity involved in the treatment was performed using the particle swarm optimization method. The execution time and the difference between the estimated parameter and the exact value were evaluated and compared with those obtained using the Levenberg-Marquardt method. The results indicate a significant agreement between the estimated and exact values in three different cases. The Levenberg-Marquardt method has a difference of 0,1942% and a performance time of 22 minutes, whereas the particle swarm optimization method has a difference of 0,0967% and a performance time of 327 minutes. The latter performs better in terms of parameter value estimation, whereas the former has better computational times. These techniques may help medical doctors to prescribe treatment protocols and may open the possibility of devising control strategies for hyperthermia therapy as a cancer treatment.

Keywords: electrical conductivity, parameter estimation, hyperthermia, Levenberg-Marquardt, radiofrequency, particle swarm optimization

RESUMEN

El uso de modelos matemáticos para el estudio de sistemas complejos como los fenómenos físicos y biológicos permite comprender su comportamiento, específicamente con respecto a variables y parámetros difíciles de obtener. Adicionalmente, el estudio de técnicas de optimización ha permitido aproximar las características de estos sistemas por medio de la correlación de simulaciones numéricas y la experimentación. La terapia de hipertermia por radiofrecuencia para el tratamiento del cáncer está actualmente en consideración para su futura aplicación médica. Sin embargo, algunas de sus propiedades son difíciles de medir, lo cual impediría su control. Este es el caso de la conductividad eléctrica, que depende de la frecuencia de inducción y de las características del tejido. En este artículo se simuló la terapia de hipertermia por radiofrecuencia mediante el método de elementos finitos. Luego se realizó una estimación de la conductividad eléctrica en el tratamiento mediante el método de optimización por enjambres de partículas. Se evaluaron el tiempo de ejecución y la diferencia del valor estimado con respecto al valor exacto, y se compararon sus valores estimados con los obtenidos mediante el método de Levenberg-Marquardt. Los resultados indican una concordancia significativa entre los valores estimados y los exactos en tres casos diferentes. El método de Levenberg-Marquardt tiene una diferencia de 0,1942% y un tiempo de ejecución de 22 minutos, mientras que el método de optimización de enjambres de partículas tiene una diferencia de 0,0967% y un tiempo de ejecución de 327 minutos. Este último tiene un mejor rendimiento en términos de estimación del valor de los parámetros, mientras que el otro tiene un mejor tiempo de ejecución computacional. Estas técnicas podrían ayudar a los médicos a prescribir protocolos de tratamiento y abrir la posibilidad de diseñar estrategias de control para la terapia de hipertermia como tratamiento para el cáncer.

Palabras clave: conductividad eléctrica, estimación de parámetros, hipertermia, Levenberg-Marquardt, radiofrecuencia, optimización por enjambre de partículas

Received: December 09th, 2020

Accepted: June 03rd, 2022

¹ Bioengineer, Universidad Santiago de Cali, Colombia. Affiliation: Young researcher, Universidad Santiago de Cali, Colombia. E-mail: jorge.lopez06@usc.edu.co

² Electronic Engineer and MSc in Electronic Engineering, Universidad del Valle, Colombia. PhD in Mechanical Engineering, Universidade Federal de Rio de Janeiro, Brazil. Affiliation: Full professor, Universidad Santiago de Cali, Colombia. E-mail: leonardo.bermeo00@usc.edu.co

How to cite: López-Pérez, J., and Bermeo, L. (2023). Estimating the Electrical Conductivity of Human Tissue in Radiofrequency Hyperthermia Therapy. *Ingeniería e Investigación*, 43(1), e92288. <http://doi.org/10.15446/ing.investig.92288>



Attribution 4.0 International (CC BY 4.0) Share - Adapt

Introduction

Mathematical modeling techniques are a tool to provide solutions to complex systems in different fields of medicine and physiology (Bratus *et al.*, 2017). In these fields, it is possible to obtain a mathematical model of any system using the principles of physics, chemistry, and biology (Selişteanu *et al.*, 2015). However, these models represent the approximate physical problem (Akhmedova and Semekin, 2013), a behavior that suggests the implementation of optimization algorithms that correlate mathematical models and experimental data in order to obtain a better understanding of the system (Rasdi *et al.*, 2016).

A considerable number of researchers use optimization algorithms such as the Levenberg-Marquardt (LM) and Particle Swarm Optimization (PSO) methods to obtain information from systems that are difficult to measure (Chen *et al.*, 2010; Pereyra *et al.*, 2013). An optimization problem consists of minimizing or maximizing an objective function, with the purpose of finding the best available information to solve a problem. In the case of a complex problem with a unique solution, an optimization algorithm might find a local minimum that does not represent the feasible solution. The PSO method avoids these local minimums, allowing to find the global optimum of the objective function (Matajira-Rueda *et al.*, 2018; Zhang, 2003).

Classical algorithms extract information sequentially, exploring the solution space in a unique direction (Chuang *et al.*, 2012), whereas optimization algorithms can explore the solution space in different directions, increasing the probability of finding a feasible solution (Cornejo and Rebolledo, 2016).

Optimization algorithms do not need to detail the structure or behavior of the system: their function is to make random changes to the probable solutions, using an adjustment function in multiple variables in order to decide which solutions are optimal and computationally efficient. The implementation of optimization algorithms allows estimating the value of a variable in order to understand the behavior of a system's parameters under different conditions.

Radiofrequency (RF) hyperthermia therapy is a treatment for the partial or total elimination of cancer cells. It consists of increasing the tissue temperature between 40 and 45 °C via the induction of radiofrequency waves. This therapy is used as adjuvant therapy in traditional cancer treatments (Colombo *et al.*, 2003; Curto, 2010; Horsman and Overgaard, 2007). Several researchers have conducted simulations of RF hyperthermia therapies in order to understand its behavior. These include the use of the finite element method (Gas, 2010; Gas and Miaskowski, 2015; Kurgan and Gas, 2009, 2010, 2011, 2015, 2016; Lv *et al.*, 2005; Miaskowski *et al.*, 2010; Miaskowski and Krawczyk, 2011; Miaskowski and Sawicki, 2013; Paruch and Turchan, 2018; Sawicki and Miaskowski, 2014; Yang *et al.*, 2005), the boundary element method (Majchrzak, Drozdek, *et al.*, 2008; Majchrzak, *et al.*,

2008; Majchrzak and Paruch, 2009, 2010), and the finite-difference method (Deng and Liu, 2002). Other works have performed state variables and parameter estimation, which correlates numerical simulations and experimental data through algorithms based on Bayesian inference.

In the study by Bermeo *et al.* (2015a), particle filters were used for the temperature field and heat source estimation. Lamien *et al.* (2017) and Bermeo *et al.* (2016a, 2016b) employed simultaneous parameters and state variables estimation while using the Liu and West filter, and Pacheco *et al.* (2020) the used Kalman filter to estimate temperature distribution. These analyses only consider a single frequency for the treatment. In the study by López *et al.* (2020), the estimation of electrical conductivity with three frequencies was performed via the LM method, with acceptable results in terms of accuracy.

Mathematical models in RF hyperthermia involve Maxwell's equations (Maxwell, 1865) and the bioheat transfer equation (Pennes, 1948). The modeling process poses a difficulty in the case of electrical conductivity in Maxwell's equations, as the values are different in intracellular and extracellular fluids, and, when the tissue is exposed to a uniform field, this behavior indicates that human tissue is inhomogeneous at a cellular scale (Ohmine *et al.*, 2004; Peters *et al.*, 2001). However, electrical conductivity can be assumed as constant in specific frequency ranges of the treatment because the differences are negligible, *i.e.*, for each frequency value, there is a value of electrical conductivity. This assumption allows the evaluation of the model under certain conditions, considering that its environment is homogeneous (Gabriel, *et al.*, 1996; Gabriel, Lau, *et al.*, 1996; Haeisen *et al.*, 1997; Kurup *et al.*, 2012; Schepps and Foster, 1980).

Thus, in this work, electrical conductivity is estimated via the PSO algorithm, and the results are compared with the estimation process performed by means of the LM method. The inverse problem was performed via simulated temperature measurements in RF hyperthermia therapy with three different frequencies and powers. The results include the analysis of the sensitivity coefficients of the system and the statistical summary of the estimation process.

Particle Swarm Optimization method (PSO)

PSO is a stochastic method for the continuous optimization of parameters based on population behavior. The algorithm imitates the interactive behavior of the flocks of birds and banks of fish to reach an optimal solution (Kennedy and Eberhart, 1995). The method solves by improving a candidate solution, updating its velocity and position according to its individual experiences, as well as those of its neighbors (Chen *et al.*, 2010). The PSO is a widely used tool for continuous optimization problems (Aazim *et al.*, 2017; Alfi, 2011; Bermeo *et al.*, 2015b; Muñoz *et al.*, 2008; J. Tang *et al.*, 2021).

In the PSO algorithm, the i -th particle is treated as a point within a space of N dimensions, and it is represented by a vector $x_{i,j}$. The best position found by the i -th particle is the position that produces the best value of the objective function, represented by $Pbest_{i,j}$, while the best position found by the entire population is represented by $Gbest_{i,j}$. The velocity for the i -th particle is represented by $V_{i,j}$. The velocity and position of the particles are calculated via Equations (1) and (2) (Kennedy and Eberhart, 1995):

$$V_{i,j}(k+1) = wV_{i,j}(k) + C_1R_1[Pbest_{i,j} - x_{i,j}(k)] + C_2R_2[Gbest_{i,j} - x_{i,j}(k)] \quad (1)$$

$$x_{i,j}(k+1) = x_{i,j}(k) + V_{i,j}(k+1) \quad (2)$$

where C_1 and C_2 are positive constants called the *acceleration coefficients*, and R_1 and R_2 are uniform random values for the interval $[0;1]$ (Lashkari and Moattar, 2016). Equation (1) calculates the new velocity at which the particle moves in the solution area as a function of the current velocity and the position. w is the inertia coefficient that influences particle velocity, which affects the exploration capability of each individual towards the best location found by it (local best) and by the swarm in the search space (global best) (Kennedy and Eberhart, 1995). It allows particles to compensate their exploration ranges between particular and global in order to locate the optimal solution within a reasonable number of iterations. Algorithm 1 shows the canonical form of the conventional PSO algorithm, where S is the objective function, Np is the number of particles population, $Pbest$ is the particle position, and $Gbest$ is the best position of the particle.

Algorithm 1. Particle swarm optimization algorithm

```

Initial randomization for particle positions and
velocities
WHILE termination criteria
FOR  $i=1$  TO  $Np$ 
  Calculate  $S_i$  of  $P_{best}$  particle
  IF  $S_i < S_{r-1}$ 
     $S_{r-1} = S_i$ 
  END IF
  Calculate the velocity of the  $P_{best}$  particle with (1)
  Calculate the position of the  $P_{best}$  particle with (2)
END FOR
  Calculate  $G_{best}$ 
END WHILE

```

Source: Authors

According to Li *et al.* (2019), the implementation of an adaptive inertia coefficient (w) allows the particle population to adjust more precisely and quickly the exploration of the optimal solution, reducing the search times for particular and global values in the conventional implementation of the PSO algorithm. On the other hand, the variation in the acceleration coefficients is used to improve the speed of convergence towards the optimal global solution, thus allowing to reach this solution efficiently. Z. Tang and Zhang

(2009) proposed a temporal variation of the acceleration coefficients in order to improve the exploration of the global optimum in the first stage of the optimization process and thus facilitate the convergence of the particles towards it. Equations (3)-(5) describe this proposal.

$$w = w_{max} - i / i_{max} (w_{max} - w_{min}) \quad (3)$$

$$C_1 = C_{min} + (C_{max} - C_{min}) / i_{th} + C_{max} \quad (4)$$

$$C_2 = C_{max} + (C_{max} - C_{min}) / i_{th} + C_{min} \quad (5)$$

The modified algorithm is shown in Algorithm 2.

Algorithm 2. Modified particle swarm optimization algorithm

```

Initial randomization for particle positions and
velocities
WHILE termination criteria
  Calculate  $w$  with (3)
  Calculate  $C_1$  with (4)
  Calculate  $C_2$  with (5)
  FOR  $i=1$  TO  $Np$ 
    Calculate  $S$  of  $P_{best}$  particle
    IF  $S_i < S_{r-1}$ 
       $S_{r-1} = S_i$ 
    END IF
    Calculate the velocity of the  $P_{best}$  particle with (1)
    Calculate the position of the  $P_{best}$  particle with (2)
  END FOR
  Calculate  $G_{best}$ 
END WHILE

```

Source: Authors

The Levenberg-Marquardt method (LM)

The LM method is used to iteratively solve nonlinear least-squares and linear problems for parameter estimation in highly complex inverse problems (Huang and Huang, 2007). This method combines the Gauss-Newton and gradient descent methods (Dattner and Gugushvili, 2018; Kanzow *et al.*, 2005) controlled by a damping factor (μ). If the damping factor tends to zero, the speed of convergence increases, and, if this factor tends to infinity, its convergence is slow and stepwise (Zhang, 2003). Equations (6) and (7) describe the parameter update and the objective function, respectively (Rouquette *et al.*, 2007).

$$P^{k+1} = P^k + [J^T J + \mu \Omega]^{-1} J^T [Y - T_{P^k}] \quad (6)$$

$$S_{k+1} = [Y - T_{P^{k+1}}]^T [Y - T_{P^{k+1}}] \quad (7)$$

where P^{k+1} is the candidate parameter, J is the sensitivity matrix, Ω is the diagonal of the matrix $J^T J$, S_{k+1} is the likelihood function, Y is the experimental temperature, and T_{P^k} is the numerical temperature. This method for estimating electrical conductivity is described in Algorithm 3.

Algorithm 3. Levenberg–Marquardt method

```

Initial randomization for the
parameter
WHILE termination criteria
Calculate  $S_{k+1}$  with (7)
Calculate  $P^{k+1}$  with (6)
Calculate
IF  $S_k < S_{k+1}$ 
     $\mu = 10 \times \mu$ 
ELSE
     $P^k = P^{k+1}$ 
     $\mu = 0,1 \times \mu$ 
END IF
END WHILE
    
```

Source: Authors

Physical problem and mathematical formulation

The physical problem considered in this work involves a cylindrical domain with the physical properties of human muscle. The heating is generated using RF through a copper coil of three spirals with a radius of 8 mm and a cross-section of 70 mm, as presented in Figure 1. The heat in the domain is due to the electrical and magnetic loss of the therapy. The boundary conditions for the material are considered to be convective heat flux in the lower and upper surfaces and adiabatic boundary conditions around the domain. The solution of the direct (forward) problem was obtained using COMSOL Multiphysics® 5.3 and verified against the results presented by Hand *et al.* (1982) and Paruch and Turchan (2018). These processes were performed on a computer with 16 GB RAM and an i7-9750H processor.

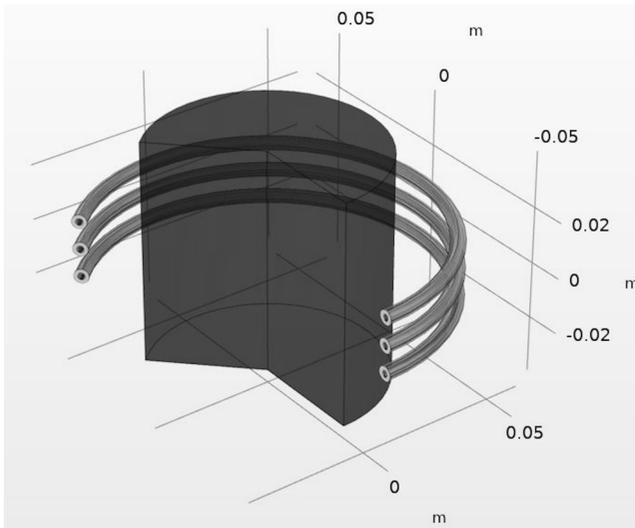


Figure 1. Domain system

Source: Authors

The electromagnetic and thermal models of the domain were obtained via Maxwell's equations (Gratiy *et al.*, 2017) and the bioheat transfer equation, respectively (Charny, 1992; Nakayama and Kuwahara, 2008; Pennes, 1948). Maxwell's equations are described in Equations (8)-(11).

$$\nabla \times \mathbf{H} = \mathbf{J} \quad (8)$$

$$\mathbf{B} = \nabla \times \mathbf{A} \quad (9)$$

$$\mathbf{J} = \sigma \mathbf{E} + \frac{\partial \mathbf{D}}{\partial t} \quad (10)$$

$$\mathbf{E} = -\frac{\partial \mathbf{A}}{\partial t} \quad (11)$$

where \mathbf{H} is the magnetic field, \mathbf{J} is the electric current density, \mathbf{B} is the magnetic flux density, \mathbf{A} is the magnetic vector potential, \mathbf{E} is the electric field, σ is the electrical conductivity, and \mathbf{D} is the electric displacement field.

The biological heat transfer is defined by Pennes (1948) as shown in Equation (12). Equation (13) presents the boundary conditions in the domain length L and the directions r, z .

$$\rho c \frac{\partial T}{\partial t} = \nabla \cdot k \nabla T - \omega_b c_b (T - T_a) + Q_m + Q \quad (12)$$

$$\left. \begin{aligned} -k \frac{\partial T}{\partial r} &= h(T - T_i) & r \geq 0 & \quad z = 0 & \quad z = L_z \\ T_{(r,z)} &= T_0 & r = L_r & \quad z \geq 0 \end{aligned} \right\} \quad (13)$$

where ρ is the density, c_b is the specific heat capacity, k is the thermal conductivity, T_0 is the initial temperature, ω_b is the blood perfusion, Q_m is the metabolic heat source, and Q is the heat source, as determined through Equations (14)-(16).

$$Q = Q_{rh} + Q_{ml} \quad (14)$$

$$Q_{rh} = \frac{1}{2} \text{Re}(\mathbf{J} \cdot \mathbf{E}') \quad (15)$$

$$Q_{ml} = \frac{1}{2} \text{Re}(i\omega \mathbf{B} \cdot \mathbf{H}') \quad (16)$$

where Q_{rh} is the electrical loss, Q_{ml} is the magnetic loss, and Re is the real part of the losses (Lakhsassasi *et al.*, 2010).

The heat source inside of the copper coil to avoid overheating is calculated by means of Equation (17).

$$Q_0 = M_t c_{pw} \frac{T_m - T}{2\pi r A_c} \quad (17)$$

where M_t is the mass flow, c_{pw} is the heat capacity of water, T_m is the initial temperature, r is the inner radius, and A_c is the cross section of the coil.

Results

In the estimation process of the electrical conductivity, the PSO and LM methods were evaluated using different

frequencies and powers (Table 1). The simulated temperature measurements were considered in the inverse problem available at the center of the domain every 30 s. To avoid an inverse crime (Kaipio and Somersalo, 2004), the simulated temperature measurements were produced on a grid of 9 128 elements, while the inverse problem solution was performed on a grid of 3 091 elements. The initial conditions of the hyperthermia model were as follows: the temperature of the environment was $T_i = 24^\circ\text{C}$, the initial temperature of the cooling pad was $T_{in} = 20^\circ\text{C}$, the convection coefficient was $h = 10 \text{ W/m}^2 \text{ K}$, the temperature in the domain was 24°C ($r = 0,04$; $-0,04 < z < 0,04$), and the thermal properties of the biological tissue Q_m , c_b , and ω_b were zero because the domain was simulated as a solid material. The parameter values were assumed constant because the effects of temperature variation on the thermal and electrical properties are negligible (Rossmann and Haemmerich, 2014). This assumption is not possible at 10,0 MHz because the temperature increase is too high. Table 1 presents the electrical properties of the phantom with three different frequencies and powers, and Table 2 shows the physical properties of the domain.

Table 1. Electric properties of the phantom (muscle properties)

Nominal frequency [MHz]	0,1	1,0	10,0
Power [W]	1 000	600	300
Electrical conductivity [S/m]	0,362	0,503	0,617
Permittivity	8 090	1 840	1 710

Source: Hasgall et al. (2018)

Table 2. Physical properties of the domain

	$k[\text{W}/(\text{m}\cdot\text{K})]$	$c_p[\text{J}/(\text{kg}\cdot\text{K})]$	$\rho[\text{kg}/\text{m}^3]$	$\sigma[\text{S}/\text{m}]$	ϵ_r	μ_r
Copper	400	385	1 090	$5,998 \times 10^7$	1	1
Air	0,024	0,240	1,086	0	1	1
Water	0,580	4,230	997	$5,500 \times 10^{-6}$	1	80
Phantom	0,49*	3 421*	1 090*	Table 1		159

Source: Comsol Multiphysics (2012) and Hasgall et al. (2018)

Temperature field

Figures 2 to 4 show the simulated temperature measurements and numerical simulation at the center of the domain $\{0,0; 0,0\}$ at 0,1, 1,0, and 10,0 MHz. These results indicate an increase in the temperature as the frequency increases. For 0,1 MHz, the temperature increase was $0,13^\circ\text{C}$; $3,23^\circ\text{C}$ for 1,0 MHz; and 45°C for 10,0 MHz. In terms of power, the temperature rises as the power increases.

To establish the appropriate frequency and power to perform RF hyperthermia, it is necessary to consider the thermal dose in the treatment. Experimental studies have revealed that patients are comfortable when the heating rate is $1,0^\circ\text{C}/\text{min}$, which does not cause considerable pain or severe damage (de Oliveira, 2014). In this study, the mean rate at 0,1 MHz was $0,02^\circ\text{C}/\text{min}$ and $0,64^\circ\text{C}/\text{min}$ at 1,0 MHz. These rates are suitable for performing mild RF hyperthermia therapy.

For the case of 10,0 MHz, the rate was $1,8^\circ\text{C}/\text{min}$, which does not allow performing therapy. However, it is possible to modify the power to reach an adequate thermal dose, e.g., a frequency of 10,0 MHz at 100 W in approximately 9 minutes (López and Bermeo, 2021).

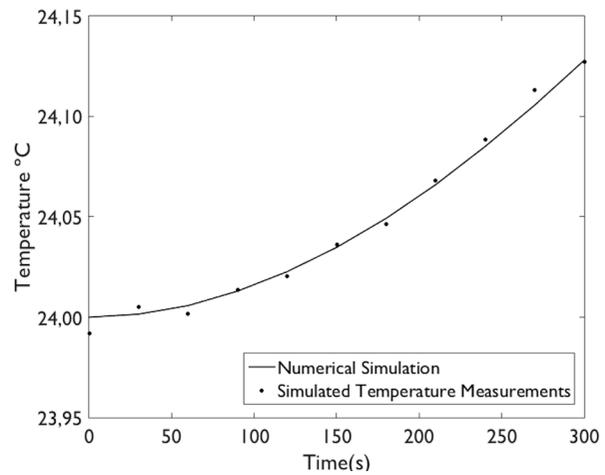


Figure 2. Temperature field at 0,1 MHz

Source: Authors

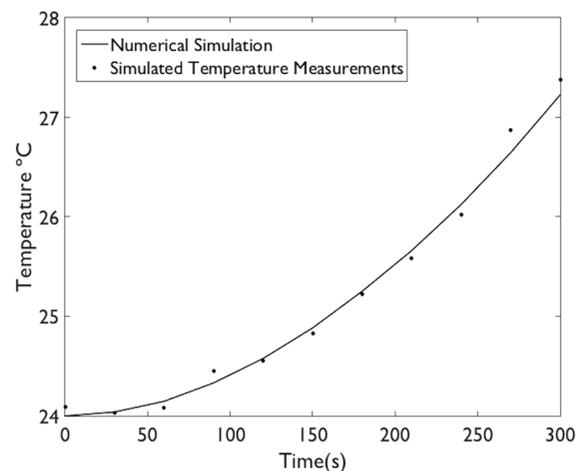


Figure 3. Temperature field at 1,0 MHz

Source: Authors

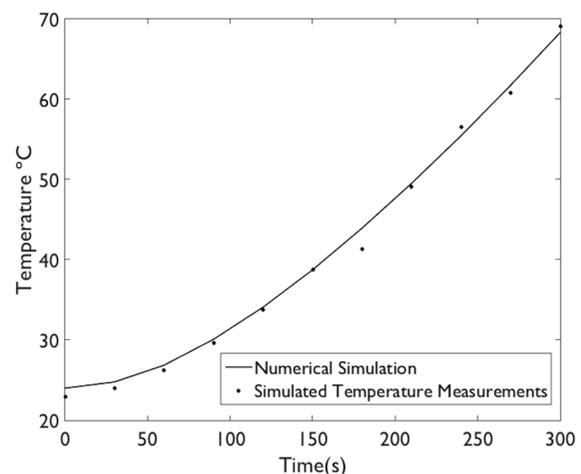


Figure 4. Temperature field at 10,0 MHz

Source: Authors

To establish the appropriate frequency and power to perform RF hyperthermia, it is necessary to consider the thermal dose in the treatment. Experimental studies have revealed that patients are comfortable when the heating rate is $1,0\text{ }^{\circ}\text{C}/\text{min}$, which does not cause considerable pain or severe damage (de Oliveira, 2014). In this study, the mean rate at $0,1\text{ MHz}$ was $0,02\text{ }^{\circ}\text{C}/\text{min}$ and $0,64\text{ }^{\circ}\text{C}/\text{min}$ at $1,0\text{ MHz}$. These rates are suitable for performing mild RF hyperthermia therapy. For the case of $10,0\text{ MHz}$, the rate was $1,8\text{ }^{\circ}\text{C}/\text{min}$, which does not allow performing therapy. However, it is possible to modify the power to reach an adequate thermal dose, e.g., a frequency of $10,0\text{ MHz}$ at 100 W in approximately 9 minutes (López and Bermeo, 2021).

Analyzing the sensitivity coefficients

The sensitivity coefficients were analyzed in order to determine the parameters that influenced the temperature field in terms of dependence and magnitude (Özisik and Orlande, 2018). The coefficients were calculated using the finite-difference approximation method, as presented in Equation (18), which indicates the changes that occur in the temperature due to a low variation in the value of the parameters.

$$J_j = \left[\frac{\partial T_i(P)}{\partial P_j} \right] \cong \frac{T_i(P_1, P_2, \dots, P_j + \varepsilon P_j, \dots, P_N) - T_i(P_1, P_2, \dots, P_j - \varepsilon P_j, \dots, P_N)}{2\varepsilon P_j} \quad (18)$$

where J is the sensitivity matrix, i is the time instant, j is the number of the parameter, and ε is the perturbation of the parameter, represented at 10% of its nominal value.

Figure 5 shows the behavior of the sensitivity coefficients of k (thermal conductivity), ρ (density), σ (electrical conductivity), μ_r (relative permeability), ε_r (relative permittivity), and c_p (heat capacity). Note that those of μ_r and ε_r have small magnitudes, indicating a low influence on the temperature field, and the remaining parameters exhibit linear dependence and large magnitudes. This behavior reveals that a single parameter can be estimated. As mentioned in this paper, electrical conductivity was estimated despite the limited information on this parameter in the literature.

Results of the estimation using the LM Method

The LM method was implemented with three different frequencies and powers (Table 1). The method was executed 50 times, and a statistical analysis was conducted in order to determine the electrical conductivity. The stopping criterion was likelihood. The study included the Shapiro-Wilk test (Shapiro and Wilk, 1965). Then, the mean, the standard deviation, and a confidence interval of 95% were calculated. Table 3 summarizes the estimation values. Note that the p-value in the Shapiro-Wilk test for the total cases is higher than 0,05, indicating that the results of 50 performances correspond to a normal distribution. In this sense, the estimation of electrical conductivity was represented by the mean, observing that the value was very close to the exact value, which indicates that the method accurately performed

the estimation. Concerning the confidence interval, note that the exact and estimated values are within this interval.

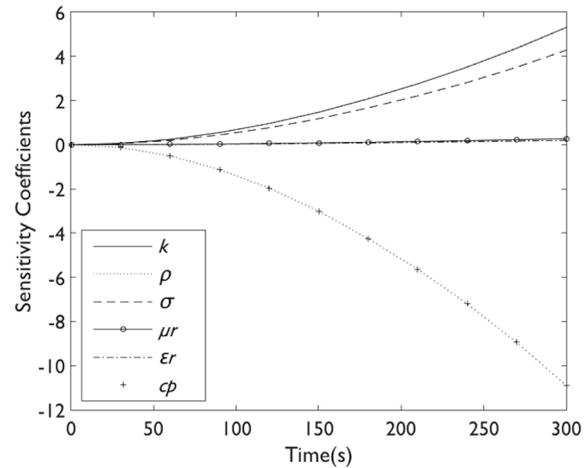


Figure 5. Sensitivity analysis
Source: Authors

Table 3. Results of the estimation of σ via the LM method

Nominal frequency (MHz)	0,1	1,0	10,0
Shapiro-Wilk test (p-value)	0,4643	0,1057	0,2615
Exact σ (S/m)	0,3620	0,5030	0,6170
Mean (S/m)	0,3604	0,5055	0,6175
Standard deviation (S/m)	0,0065	0,0130	0,0013
Confidence interval (S/m)	0,3513-0,3641	0,5009-0,5102	0,6170-0,6210
Computational time (min)	20,0856	22,1876	23,9812

Source: Authors

Results of the estimation using the PSO Method

The PSO method was implemented using the same condition as the LM method. The algorithm was executed 50 times with a population of 50 particles for every computer simulation. The stopping criterion was the maximum number of iterations (50 by default). The study included the Shapiro-Wilk normality test, and the results are presented in Table 4. Note that the results of the 50 performances correspond to a normal distribution. Thus, the mean of a global best was very close to the exact value. The results of estimates are similar to those found by the LM method.

Discussions

Tables 3 and 4 show the statistical analysis for the methods implemented in RF hyperthermia therapy, demonstrating that the results obtained from the estimation are close to the exact value of electrical conductivity, with a difference of

0,1942%. Note that the maximum standard deviation of the total cases was 0,0130 S/m, indicating that the estimation values have good precision. This behavior was verified with the 95% confidence interval, which reveals that, within the range obtained, the exact value of the parameter lies within 95% certainty.

Table 4. Results of the estimation of σ via PSO

Nominal frequency (MHz)	0,1	1,0	10,0
Shapiro Wilk test (p-value)	0,0764	0,2177	0,1432
Exact σ (S/m)	0,3620	0,5030	0,6170
Mean (S/m)	0,3614	0,5033	0,6166
Standard deviation (S/m)	0,0021	0,0009	0,0016
Confidence interval (S/m)	0,3606-0,3621	0,5031-0,5036	0,6160-0,6171
Computational time (min)	315,1985	380,3298	286,1487

Source: Authors

In the case of the LM method, 50 estimations were performed. Note that the mean values were very close to the estimated parameter. The estimated values were as follows: 0,3604, 0,5055, and 0,6175 S/m for 0,1, 1,0, and 10,0 MHz, respectively. These results indicate that the algorithm has a good convergence to the exact value. In terms of the computational time, the mean time of each estimation process was approximately 22 minutes, indicating that it is a fast-convergence algorithm. The PSO method was evaluated on a population of 50 particles and 50 iterations as a stopping criterion. Also, 50 estimation processes were performed, where the solution was obtained from the position of the best particle. The estimated values were 0,3614, 0,5033, and 0,6166 S/m for 0,1, 1,0, and 10,0 MHz, respectively. Note that this algorithm achieved an excellent convergence of the electrical conductivity value, with a difference of 0,0967%. In terms of the computational time, the mean time of each estimation process was approximately 327 minutes, which indicates that it is an acceptable convergence algorithm. This time could be lower if another stopping criterion were considered instead of the number of iterations, (e.g., the mean square error).

These results aim to predict the electrical conductivity in biological tissues using mathematical simulations and optimization algorithms on phantoms, where the final goal is to improve RF hyperthermia in terms of frequency and exposure time values, which may help medical doctors with the planning of individual treatment protocols. This also opens the possibility to design real-time control strategies.

Conclusions

This paper presents the application of two methods for estimating electrical conductivity in RF hyperthermia

therapy for cancer treatment. The estimation problem was performed using the LM and PSO methods on a rectangular 2D axisymmetric system with temperature measurements available at one single location inside the domain. The estimation process was executed with three different frequencies and powers. The results were more accurate and smoother with the PSO method, and an excellent agreement with the exact values was obtained. In terms of computational time, the LM method performed better because its stopping criterion depends on the likelihood, whereas, for the PSO method, it is the number of iterations.

Acknowledgements

The authors are thankful for the support provided by DGI of Universidad Santiago de Cali, Colombia, project No. 819-621120-1767 and young researcher grant No. 05-2021.

References

- Aazim, R., Liu, C., Haaris, R., and Mansoor, A. (2017). Rapid generation of control parameters of multi-infeed system through online simulation. *Ingeniería e Investigación*, 37(2), 67-73. <https://doi.org/10.15446/ing.investig.v37n2.61822>
- Akhmedova, S., and Semenkina, E. (2013). Co-operation of biology related algorithms. In IEEE (Eds.), *2013 IEEE Congress on Evolutionary Computation, CEC 2013* (pp. 2207-2214). IEEE. <https://doi.org/10.1109/CEC.2013.6557831>
- Alfi, A. (2011). PSO with adaptive mutation and inertia weight and its application in parameter estimation of dynamic systems. *Acta Automatica Sinica*, 37(5), 541-549. [https://doi.org/10.1016/s1874-1029\(11\)60205-x](https://doi.org/10.1016/s1874-1029(11)60205-x)
- Bermeo, L. A., Caicedo, E., Clementi, L., and Vega, J. (2015). Estimation of the particle size distribution of colloids from multiangle dynamic light scattering measurements with particle swarm optimization. *Ingeniería e Investigación*, 35(1), 49-54. <https://doi.org/10.15446/ing.investig.v35n1.45213>
- Bermeo, L. A., Orlande, H. R. B., and Elicabe, G. E. (2015). Estimation of state variables in the hyperthermia therapy of cancer with heating imposed by radiofrequency electromagnetic waves. *International Journal of Thermal Sciences*, 98, 228-236. <https://doi.org/10.1016/j.ijthermalsci.2015.06.022>
- Bermeo, L. A., Orlande, H. R. B., and Elicabe, G. E. (2016a). Combined parameter and state estimation in the radiofrequency hyperthermia treatment of cancer. *Heat Transfer, Part A: Applications*, 70(6), 581-594. <https://doi.org/10.1080/10407782.2016.1193342>
- Bermeo, L. A., Orlande, H. R. B., and Elicabe, G. E. (2016b). Combined parameter and state estimation problem in a complex domain: RF hyperthermia treatment using nanoparticles. *Journal of Physics: Conferences Series*, 745(032014), 1-8. <https://doi.org/10.1088/1742-6596/745/3/032014>
- Bratus, A., Samokhin, I., Yegorov, I., and Yurchenko, D. (2017). Maximization of viability time in a mathematical model of cancer therapy. *Mathematical Biosciences*, 294, 110-119. <https://doi.org/10.1016/j.mbs.2017.10.011>

- Charny, C. K. (1992). Mathematical models of bioheat transfer. *Advances in Heat Transfer*, 22(C), 19-155. [https://doi.org/10.1016/S0065-2717\(08\)70344-7](https://doi.org/10.1016/S0065-2717(08)70344-7)
- Chen, W. N., Zhang, J., Chung, H. S. H., Zhong, W. L., Wu, W. G., and Shi, Y. H. (2010). A novel set-based particle swarm optimization method for discrete optimization problems. *IEEE Transactions on Evolutionary Computation*, 14(2), 278-300. <https://doi.org/10.1109/TEVC.2009.2030331>
- Chuang, L. Y., Lin, Y. Da, Chang, H. W., and Yang, C. H. (2012). An improved PSO algorithm for generating protective SNP barcodes in breast cancer. *PLoS ONE*, 7(5), 0037018. <https://doi.org/10.1371/journal.pone.0037018>
- Colombo, R., da Pozzo, L. F., Salonia, A., Rigatti, P., Leib, Z., Baniel, J., Caldarera, E., and Pavone-Macaluso, M. (2003). Multicentric study comparing intravesical chemotherapy alone and with local microwave hyperthermia for prophylaxis of recurrence of superficial transitional cell carcinoma. *Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology*, 21(23), 4270-4276. <https://doi.org/10.1200/JCO.2003.01.089>
- Comsol Multiphysics (2012). *The RF module user's guide*. <https://doc.comsol.com/5.3/doc/com.comsol.help.rf/RFModuleUsersGuide.pdf>
- Cornejo, O., and Rebolledo, R. (2016). Estimación de parámetros en modelos no lineales: algoritmos y aplicaciones. *Revista EIA*, 13(25), 81-98. <https://doi.org/10.14508/reia.2016.13.25.81-98>
- Curto, S. (2010). *Antenna development for radio frequency hyperthermia applications* [Doctoral thesis, Dublin Institute of Technology]. <https://doi.org/10.21427/D7CP65>
- Dattner, I., and Gugushvili, S. (2018). Application of one-step method to parameter estimation in ODE models. *Statistica Neerlandica*, 72(2), 126-156. <https://doi.org/10.1111/stan.12124>
- Deng, Z.-S., and Liu, J. (2002). Monte Carlo method to solve multidimensional bioheat transfer problem. *Numerical Heat Transfer, Part B*, 42, 543-567. <https://doi.org/10.1080/10407790190054076>
- Gabriel, S., Gabriel, C., and Corthout, E. (1996). The dielectric properties of biological tissues: I. Literature survey. *Physics in Medicine and Biology*, 41(11), 2231-2249. <https://doi.org/10.1088/0031-9155/41/11/001>
- Gabriel, S., Lau, R. W., and Gabriel, C. (1996). The dielectric properties of biological tissues: III. Parametric models for the dielectric spectrum of tissues. *Physics in Medicine and Biology*, 41(11), 2271-2293. <https://doi.org/10.1088/0031-9155/41/11/003>
- Gas, P. (2010). Temperature inside tumor as time function in RF hyperthermia. *Przeegląd Elektrotechniczny*, 86(12), 42-45.
- Gas, P., and Miaskowski, A. (2015, September 17-19). *Specifying the ferrofluid parameters important from the viewpoint of Magnetic Fluid Hyperthermia* [Conference presentation]. 2015 Selected Problems of Electrical Engineering and Electronics (WZEE), Kielce, Poland. <https://doi.org/10.1109/WZEE.2015.7394040>
- Gratiy, S. L., Haines, G., Denman, D., Hawrylycz, M. J., Koch, C., Einevoll, G. T., and Anastassiou, C. A. (2017). From Maxwell's equations to the theory of current-source density analysis. *European Journal of Neuroscience*, 45(8), 1013-1023. <https://doi.org/10.1111/ejn.13534>
- Hand, J. W., Ledda, J. L., and Evans, N. T. S. (1982). Considerations of radiofrequency induction heating for localised hyperthermia. *Physics in Medicine and Biology*, 27(1), 1-16. <https://doi.org/10.1088/0031-9155/27/1/001>
- Hasgall, P. A., di Gennaro, F., Baumgartner, C., Neufeld, E., Lloyd, B., Gosselin, M., Payne, D., Klingensböck, A., and Kuster, N. (2018). *IT'IS Database for thermal and electromagnetic parameters of biological tissues, Version 4.0*. <https://doi.org/10.13099/VIP21000-04-0>
- Hauelsen, J., Ramon, C., Eiselt, M., Brauer, H., and Nowak, H. (1997). Influence of tissue resistivities on neuromagnetic fields and electric potentials studied with a finite element model of the head. *IEEE Transactions on Biomedical Engineering*, 44(8), 727-735. <https://doi.org/10.1109/10.605429>
- Horsman, M. R., and Overgaard, J. (2007). Hyperthermia: A potent enhancer of radiotherapy. *Clinical Oncology*, 19, 418-426. <https://doi.org/10.1016/j.clon.2007.03.015>
- Huang, C. H., and Huang, C. Y. (2007). An inverse problem in estimating simultaneously the effective thermal conductivity and volumetric heat capacity of biological tissue. *Applied Mathematical Modelling*, 31(9), 1785-1797. <https://doi.org/10.1016/j.apm.2006.06.002>
- Kaipio, J. P., and Somersalo, E. (2004). *Computational and statistical methods for inverse problems*. Springer.
- Kanzow, C., Yamashita, N., and Fukushima, M. (2005). Levenberg-Marquardt methods with strong local convergence properties for solving nonlinear equations with convex constraints. *Journal of Computational and Applied Mathematics*, 173(2), 321-343. <https://doi.org/10.1016/j.cam.2004.03.015>
- Kennedy, J., and Eberhart, R. (1995). Particle swarm optimization. *Proceedings of ICNN'95 - International Conference on Neural Networks*, 4(2), 1942-1948. <https://doi.org/10.1109/ICNN.1995.488968>
- Kurgan, E., and Gas, P. (2009). Distribution of the temperature in human body in RF hyperthermia. *Przeegląd Elektrotechniczny*, 85(12), 96-99.
- Kurgan, E., and Gas, P. (2010). Estimation of temperature distribution inside tissues in external RF hyperthermia. *Przeegląd Elektrotechniczny*, 86(01), 100-102.
- Kurgan, E., and Gas, P. (2011). Treatment of tumors located in the human thigh using RF hyperthermia. *Przeegląd Elektrotechniczny*, 87(12), 103-106.
- Kurgan, E., and Gas, P. (2015). Simulation of the electromagnetic field and temperature distribution in human tissue in RF hyperthermia. *Przeegląd Elektrotechniczny*, 91(1), 169-172. <https://doi.org/10.15199/48.2015.01.37>
- Kurgan, E., and Gas, P. (2016, September 14-17). *Analysis of electromagnetic heating in magnetic fluid deep hyperthermia* [Conference presentation]. 7th International Conference Computational Problems of Electrical Engineering (CPEE), Sandomierz, Poland. <https://doi.org/10.1109/CPEE.2016.7738756>
- Kurup, D., Joseph, W., Vermeeren, G., and Martens, L. (2012). In-body path loss model for homogeneous human tissues. *IEEE Transactions on Electromagnetic Compatibility*, 54(3), 556-564. <https://doi.org/10.1109/TEM.2011.2164803>

- Lakhssassi, A., Kengne, E., and Semmaoui, H. (2010). Modified Pennes' equation modelling bio-heat transfer in living tissues: analytical and numerical analysis. *Natural Science*, 02(12), 1375-1385. <https://doi.org/10.4236/ns.2010.212168>
- Lamien, B., Bermeo, L. A., Orlande, H. R. B., and Eliçabe, G. E. (2017). State estimation in bioheat transfer: A comparison of particle filter algorithms. *International Journal of Numerical Methods for Heat & Fluid Flow*, 27(3), 615-638. <https://doi.org/10.1108/HFF-03-2016-0118>
- Lashkari, M., and Moattar, M. H. (2016, November 11-12). *The improved K-means clustering algorithm using the proposed extended PSO algorithm* [Conference presentation]. 2015 International Congress on Technology, Communication and Knowledge (ICTCK), Mashhad, Iran. <https://doi.org/10.1109/ICTCK.2015.7582708>
- Li, C., Liu, C., Yang, L., He, L., and Wu, T. (2019). Particle swarm optimization for positioning the coil of transcranial magnetic stimulation. *BioMed Research International*, 2019, 9461018. <https://doi.org/10.1155/2019/9461018>
- López, J. I., and Bermeo, L. A. (2021). Parametric study of thermal damage in the hyperthermia treatment by radiofrequency. *2021 IEEE 2nd International Congress of Biomedical Engineering and Bioengineering (CI-IB&BI)*, 7, 1-4. <https://doi.org/10.1109/CI-IB&BI54220.2021.9626117>
- López, J. I., Serna, R. D., Bermeo, L. A., and Castillo, J. F. (2020). Estimation of electrical conductivity from radiofrequency hyperthermia therapy for cancer treatment by Levenberg Marquardt method. *Communications in Computer and Information Science*, 1195, 141-152. https://doi.org/10.1007/978-3-030-42531-9_12
- Lv, Y. G., Deng, Z. S., and Liu, J. (2005). 3-D Numerical study on the induced heating effects of embedded micro/nanoparticles on human body subject to external medical electromagnetic field. *IEEE Transactions on Nanobioscience*, 4(4), 284-294. <https://doi.org/10.1109/TNB.2005.859549>
- Majchrzak, E., Drozdek, J., and Paruch, M. (2008). Heating of tissue by means of the electric field: Numerical model basing on the BEM. *Scientific Research of the Institute of Mathematics and Computer Science*, 7(1), 99-110.
- Majchrzak, E., Dziatkiewicz, G., and Paruch, M. (2008). The modelling of heating a tissue subjected to external electromagnetic field. *Acta of Bioengineering and Biomechanics/Wroc aw University of Technology*, 10(2), 29-37.
- Majchrzak, E., and Paruch, M. (2009). Numerical modelling of temperature field in the tissue with a tumor subjected to the action of two external electrodes. *Scientific Research of the Institute of Mathematics and Computer Science*, 8(1), 137-145.
- Majchrzak, E., and Paruch, M. (2010). Numerical modelling of tissue heating by means of the electromagnetic field. *Scientific Research of the Institute of Mathematics and Computer Science*, 9(1), 89-97.
- Matajira-Rueda, D., Cruz-Duarte, J., Aviña-Cervantes, J., and Correa-Cely, C. (2018). Global optimization algorithms applied in a parameter estimation strategy. *Revista UIS Ingenierías*, 17(1), 233-242. <https://doi.org/10.18273/revuin.v17n1-2018023>
- Maxwell, J. C. (1865). A dynamical theory of the electromagnetic field. *Philosophical Transactions of the Royal Society of London*, 155, 459-512. <https://doi.org/10.5479/sil.423156.39088007130693>
- Miaskowski, A., and Krawczyk, A. (2011). Magnetic fluid hyperthermia for cancer therapy. *Przegląd Elektrotechniczny*, 87(12), 125-127.
- Miaskowski, A., and Sawicki, B. (2013). Magnetic fluid hyperthermia modeling based on phantom measurements and realistic breast model. *IEEE Transactions on Biomedical Engineering*, 60(7), 1806-1813. <https://doi.org/10.1109/TBME.2013.2242071>
- Miaskowski, A., Sawicki, B., Krawczyk, A., and Yamada, S. (2010). The application of magnetic fluid hyperthermia to breast cancer treatment. *Przegląd Elektrotechniczny*, 86(12), 99-101.
- Muñoz, M. A., López, J. A., and Caicedo, E. F. (2008). Swarm intelligence problem-solving societies (a review). *Ingeniería e Investigación*, 28(2), 119-130. <https://doi.org/10.15446/ing.investig.v28n2.14901>
- Nakayama, A., and Kuwahara, F. (2008). A general bioheat transfer model based on the theory of porous media. *International Journal of Heat and Mass Transfer*, 51(11-12), 3190-3199. <https://doi.org/10.1016/j.ijheatmasstransfer.2007.05.030>
- Ohmine, Y., Morimoto, T., Kinouchi, Y., Iritani, T., Takeuchi, M., Haku, M., and Nishitani, H. (2004). Basic study of new diagnostic modality according to noninvasive measurement of the electrical conductivity of tissues. *The Journal of Medical Investigation*, 51(3,4), 218-225. <https://doi.org/10.2152/jmi.51.218>
- Özisik, M. N., and Orlande, H. R. B. (2018). *Inverse heat transfer: Fundamentals and applications*. Routledge. <https://doi.org/10.1201/9780203749784>
- Pacheco, C. C., Orlande, H. R. B., Colaço, M. J., Dulikravich, G. S., Varon, L. A. B., and Lamien, B. (2020). Real-time temperature estimation with enhanced spatial resolution during MR-guided hyperthermia therapy. *Numerical Heat Transfer, Part A: Applications*, 77(8), 782-806. <https://doi.org/10.1080/01407782.2020.1720409>
- Paruch, M., and Turchan, Ł. (2018). Mathematical modelling of the destruction degree of cancer under the influence of a RF hyperthermia. *AIP Conference Proceedings*, 1922, 060003. <https://doi.org/10.1063/1.5019064>
- Pennes, H. H. (1948). Analysis of tissue and arterial blood temperatures. *Journal of Applied Physiology*, 1(2), 93-122. <https://doi.org/10.1152/jappl.1948.1.2.93>
- Pereyra, S., Lombera, G. A., Frontini, G., and Urquiza, S. A. (2013). Sensitivity analysis and parameter estimation of heat transfer and material flow models in friction stir welding. *Materials Research*, 17(2), 397-404. <https://doi.org/10.1590/s1516-14392013005000184>
- Peters, M. J., Stinstra, J. G., and Hendriks, M. (2001). Estimation of the electrical conductivity of human tissue. *Electromagnetics*, 21(7-8), 545-557. <https://doi.org/10.1080/027263401752246199>
- Rasdi, L. M., Fanany, M. I., and Arymurthy, A. M. (2016). Meta-heuristic algorithms for convolution neural network. *Computational Intelligence and Neuroscience*, 2016, 1537325. <https://doi.org/10.1155/2016/1537325>

- Rossmann, C., and Haemmerich, D. (2014). Review of temperature dependence of thermal properties, dielectric properties, and perfusion of biological tissues at hyperthermic and ablation temperatures. *Critical Reviews in Biomedical Engineering*, 42(6), 467-492. <https://doi.org/10.1615/critrevbio-medeng.2015012486>
- Rouquette, S., Guo, J., and Le Masson, P. (2007). Estimation of the parameters of a Gaussian heat source by the Levenberg–Marquardt method: Application to the electron beam welding. *International Journal of Thermal Sciences*, 46(2), 128-138. <https://doi.org/10.1016/j.ijthermalsci.2006.04.015>
- Sawicki, B., and Miaskowski, A. (2014). Nonlinear higher-order transient solver for magnetic fluid hyperthermia. *Journal of Computational and Applied Mathematics*, 270, 143-151. <https://doi.org/10.1016/j.cam.2014.02.008>
- Schepps, J. L., and Foster, K. R. (1980). The UHF and microwave dielectric properties of normal and tumour tissues: Variation in dielectric properties with tissue water content. *Physics in Medicine and Biology*, 25(6), 1149-1159. <https://doi.org/10.1088/0031-9155/25/6/012>
- Selișteanu, D., Endrescu, D., Georgeanu, V., and Roman, M. (2015). Mammalian cell culture process for monoclonal antibody production: Nonlinear modelling and parameter estimation. *BioMed Research International*, 2015, 598721. <https://doi.org/10.1155/2015/598721>
- Shapiro, S. S., & Wilk, M. B. (1965). An Analysis of Variance Test for Normality (Complete Samples). *Biometrika*, 52(3/4), 591. <https://doi.org/10.2307/2333709>
- Tang, J., Liu, G., and Pan, Q. (2021). A review on representative swarm intelligence algorithms for solving optimization problems: Applications and trends. *IEEE/CAA Journal of Automatica Sinica*, 8(10), 1627-1643. <https://doi.org/10.1109/JAS.2021.1004129>
- Tang, Z., and Zhang, D. (2009). A modified particle swarm optimization with an adaptive acceleration coefficients. *Proceedings - 2009 Asia-Pacific Conference on Information Processing, APCIP 2009*, 2, 330-332. <https://doi.org/10.1109/APCIP.2009.217>
- Yang, X., Du, J., and Liu, Y. (2005). Advances in hyperthermia technology. *Proceedings of the 2005 IEEE Engineering in Medicine and Biology 27th Annual Conference*, 7, 6766-6769. <https://doi.org/10.1109/IEMBS.2005.1616058>
- Zhang, J.-L. (2003). On the convergence properties of the Levenberg–Marquardt method. *Optimization*, 52(6), 739-756. <https://doi.org/10.1080/0233193031000163993>