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A 3D finite element model for hyperthermia injury of blood-perfused skin

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Abstract

One of the objectives of this study is to propose a numerical model of thermal damage to the skin. This model simulates the propagation of a burn and suggests treatments to prevent it from spreading. In order to achieve this goal, we developed a 3D multi-layer finite elements model of the skin coupled to a model presenting hyperthermic damage. The numerical model of the skin not only takes account of the various layers’ thermal properties, but also of blood perfusion and veins. The model of damage is based on a law of Arrhenius. We tested two various quick intervention treatments so as to prevent the burn from spreading. The first treatment consists in cooling the burned zone with a flow of cool water at 10°C, while the second solution simulates the apposition of ice on the burn. The results show that according to the severity of the burn the second treatment seems to be the most appropriate. Moreover, our model opens interesting prospects in the analysis of hyperthermic damage.

Thermal damage; Skin; Finite element; Burn limitation

1 Introduction

Hyperthermia injury occurs after temperature elevation in tissues over a threshold value for a period of time. The first relationship between temperature, exposure duration and some specific degree of thermal injury is due to (Moritz and Henriques, 1947a), (Moritz and Henriques, 1947b). The damage produced was modeled by an Arrhenius equation type where a rate of chemical process is supposed to depend on the exposure duration for a given temperature. Twelve years later (Stoll and Greene, 1959) proposed two kinetic equations:

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one for low temperatures and the other for high temperatures. The parameters of the damage law were different from those chosen by Henriques and Moritz. Since these publications, the scientific community worked to determine the parameters of the damage law. In his paper (Wright, 2003), N. T. Wright presented a large review of Arrhenius’s parameters, found in literature and underlined a relationship between them.

In parallel, the scientific community worked to build predictive numerical models of hyperthermia injury. In 1983, Diller and Hayes (Diller and Hayes, 1983) presented a finite element model of burn injuries in blood-perfused skin. Since this date, many models and studies were carried out on the subject (Jiang et al., 2002), (Ng and Chua, 2002), (Chang and Nguyen, 2004), (Sramek et al., 2009), (Fuentes et al., 2010). Different sources of burns (laser, radio frequency, fluid, ...) were simulated on different tissues (skin, retinal, liver, ...). Among these publications, recent numerical approaches (Fuentes et al., 2010), (Museux et al., 2012) gave good predictive results on animal tissues.

The numerical model of thermal damage developed here aims at simulating the propagation of a skin burn and at proposing treatments to limit its expansion. Similar studies were presented by (Ng and Chua, 2002) and by (Becker and Kuznetsov, 2007). Here, we propose a 3D multi-layer finite elements model of the skin coupled to a model of hyperthermic damage. The numerical model of the skin not only takes account of the various layers thermal properties, but also of blood perfusion and veins. It was validated using infrared measurements on human subjects in (Ratovoson et al., 2011). The model of damage is based on a law of Arrhenius. We propose to simulate two treatments of quick intervention in order to prevent the burn from spreading. The first treatment consists in cooling the burned zone thanks to a flow of cool water, while the second method simulates the apposition of ice on the burn.

2 Modeling

The model proposed in this paper, is a three-dimensional multi-layer in which the veins are considered as fluid continuous media. As reported in (Becker and Kuznetsov, 2007) the skin is composed of 4 layers, which are: the stratum corneum, the epidermis, the dermis and the hypodermis. They highlighted the importance to consider composite structures for skin modeling. As in papers (Jiang et al., 2002), (Museux et al., 2012), (Becker and Kuznetsov, 2007), (Becker and Kuznetsov, 2007) the epidermis, the dermis and the hypodermis are solid continuous media in which the heat transfer is controlled by a Pennes equation type. To complement the numerical model, the thermal damage is controled by an Arrhenius law.

2.1 Heat transfer in a fluid continuous medium

To model blood, the equations governing heat transfer in a fluid continuous medium are presented. The velocity vector of a particle $P$ of position $(x, y, z)$ at time $t$ is noted $U = (u_x, u_y, u_z)$. The strain rate tensor is defined by

$$
\varepsilon = \frac{1}{2}(\nabla u + \nabla u^T)
$$

The temperature is noted as $T$. Heat transfer in a homogeneous, incompressible and
viscous fluid is controlled by the following partial derivative equation

$$\rho C \frac{\partial T}{\partial t} - k \Delta T + \rho C U \cdot \nabla T = \mu (2\varepsilon - \frac{2}{3} \nabla \cdot u) : \nabla u + r$$  \hspace{1cm} (1)

where $\rho$ is the specific mass (kg.m$^{-3}$),
$C$ is the specific heat (J.kg$^{-1}$.oC$^{-1}$),
$k$ is the thermal conductivity (W.m$^{-1}$.oC$^{-1}$),
$\mu$ is the viscosity (Pa.s),
$r$ is the heat source (W.m$^{-3}$). (neglected in our study)

2.2 Pennes equation

To model the biological tissues constituting the skin, the Pennes model was chosen. The first biothermal equation was developed by Pennes in 1948 (Pennes, 1948). There are two mechanisms at stake when we deal with heat transfer in living bodies: blood flow and metabolism. Before undertaking any study, it is necessary to determine which scale to consider in studying heat transfers. Indeed the size of the blood system may vary. The Pennes model proves to be the most appropriate and effective, with a scale ranging from a millimeter to several centimeters (Weinbaum and Jiji, 1985). Pennes made the assumption that in a given volume of study, heat transfer between the blood and the tissue is proportional to the temperature difference between the arterial blood temperature $T_a$ (entering the volume of study) and the (outgoing) venous blood temperature $T_v$. The proportionality factor $w$ is called rate of perfusion. The metabolism is also included in the biothermal equation as a distribution of energy source:

$$Q_{met} = Q_{met}^o Q_{10}^{T_a - T_v}$$  \hspace{1cm} (2)

where $Q_{met}^o$ is the basal metabolism at 37$^\circ$C, $Q_{10}$ is the coefficient of temperature dependence ranging between 2 and 3 W.m$^{-3}$ and $T_o$ is chosen at 37$^\circ$C.

The general biothermal equation obtained with the balance of energy for a volume of tissue taking account of the blood perfusion and the metabolism is given by:

$$\rho C \frac{\partial T}{\partial t} - \nabla k_t \cdot \nabla T = w \rho_b C_b (T_a - T_v) + Q_{met}$$  \hspace{1cm} (3)

where $\rho C \frac{\partial T}{\partial t}$ is the stored energy (W.m$^{-3}$), $\nabla k_t \cdot \nabla T$ is the heat conduction (W.m$^{-3}$), $k_t$ is the thermal conductivity of the tissue, $w(\rho C)_b (T_a - T_v)$ is the heat generated by perfusion (W.m$^{-3}$). The index $b$ is reserved for blood coefficient of materials. $Q_{met}$ is the metabolic internal power therefore generated (W.m$^{-3}$).

For years, the Pennes model was criticized by some and defended by others. Despite these controversies and criticisms, most mathematical analyses carried out in biothermal transfers continue to be based on this equation (Klinger, 1974), (Weinbaum et al., 1984), (Jiang et al., 2002), (Ng and Chua, 2002), (Sramek et al., 2009), (Museux et al., 2012).
2.3 Assumptions

To solve the problem, the following assumptions were made:

- Only two layers (dermis and hypodermis) were taken into account. The stratum corneum and the epidermis were neglected because of their lack of thickness.
- \( k_t \) was chosen constant, thus \( \nabla k_t \nabla T = k_t \Delta T \).
- \( T_v = T \)
- The generation of metabolic internal power \( Q_{met} \) was taken as constant in the dermis and the hypodermis.
- A model of 10x4 cm\(^2\) of surface and 4 mm of thickness was used (see figure 1).

![Figure 1: 3D model of skin](image)

- The vein is placed between the dermis and the hypodermis. Blood circulation was created by using a material having transport velocity properties: equation [1].
- The position of the vein is determined by the coordinates of its axis\(\hat{s}\)s center \( x_c \) and \( y_c \).
- The blood is assumed to be an incompressible and viscous fluid and the flow is assumed to be stationary. The velocity of the blood is then a parabolic function of direction \( \vec{z} \):

\[
    u_z(x, y, z) = -\frac{U_{max}}{R^2}[(x - x_c)^2 + (y - y_c)^2] + U_{max}
\]

where \( U_{max} \) is the maximum velocity in the section of the vein and \( R \) is the radius of the
vein. According to (Ratovoson et al., 2011), $U_{max}$ was taken at a value equals to 2.3 cm/s and vein depth was 1 mm.

- The properties of materials are coming from (Gowrishankar et al., 2004) and collected in table 1.

<table>
<thead>
<tr>
<th>Sub-domain number</th>
<th>Layer</th>
<th>Thickness (m)</th>
<th>Specific heat $C$ (J.kg$^{-1}$.°C$^{-1}$)</th>
<th>Thermal conductivity $k$ (W.m$^{-1}$.°C$^{-1}$)</th>
<th>Specific mass $\rho$ (kg.m$^{-3}$)</th>
<th>Rate of perfusion $w$ (s$^{-1}$)</th>
</tr>
</thead>
<tbody>
<tr>
<td>0</td>
<td>Epidermis</td>
<td>0.08 $10^{-3}$</td>
<td>3590</td>
<td>0.24</td>
<td>1200</td>
<td>0</td>
</tr>
<tr>
<td>1</td>
<td>Dermis</td>
<td>0.001</td>
<td>3300</td>
<td>0.45</td>
<td>1200</td>
<td>0.00125</td>
</tr>
<tr>
<td>2</td>
<td>Blood</td>
<td>0.0013*</td>
<td>3770</td>
<td>0.45</td>
<td>1060</td>
<td>0</td>
</tr>
<tr>
<td>3</td>
<td>Hypodermis</td>
<td>0.003</td>
<td>2500</td>
<td>0.19</td>
<td>1000</td>
<td>0.00125</td>
</tr>
</tbody>
</table>

Table 1: Material properties: 0.0013* is the diameter of the veins and can vary from 1 to 3 mm

All these assumptions and parameter values have been validated for a forearm skin using infrared and echo Doppler measurements (see our previous paper (Ratovoson et al., 2011)).

### 2.4 Damage model

The rate of thermal damage in tissues is used to evaluate and quantify the damage undergone by the skin following a burn. One can evaluate the rate of thermal damage by:

\[
\frac{d\Omega}{dt} = Ae^{-\frac{E_a}{RT(t)}}
\]  

(4)

where $\Omega$ stands for the thermal damage and is uniteless, $A$ is the frequency factor (expressed in s$^{-1}$). It can take values from $4.11 \times 10^{53}$ to $9.39 \times 10^{104}$ [s$^{-1}$] (Wright, 2003) for epidermis tissues. $E_a$ is the activation energy for skin and is varying from $3.39 \times 10^{8}$ to $6.65 \times 10^{8}$ [J.kmol$^{-1}$]. $R = 8.315$ J.mol$^{-1}.K^{-1}$, is the universal gas constant.

### 2.5 Initial and boundary conditions

- Permanent boundary conditions: A temperature $T_a$ was imposed on the lower part of the hypodermis, on the lateral walls and at the entrance of the vein (scheme (a) of figure 2). On the front and back surfaces heat flux was null (scheme (b) of figure 2). The upper part of the dermis (except the disc of radius : 1 cm) was subject to heat transfer by convection (scheme (c) of figure 2). Convection was modeled by Newton’s law of cooling:

\[
k\frac{\partial T}{\partial n} = h(T - T_e)
\]

where $n$ is the normal vector at the surface, $h$ is the coefficient of convection and $T_e$ is the temperature of air. We chose $h = 7$ W.m$^{-2}.°C^{-1}$ (see (Xu et al., 2008)) and $T_e = 20°C$.

The connection between the layers was ensured by the continuity of the heat flow.
- In order to ensure a stationary distribution of temperature in tissues, before the burning process, an initialization step of sufficient duration (200 seconds) was included. Then, simulations of burn and health care treatments were carried out in three steps. On the disc of 1 cm of radius on the dermis’ upper part (scheme (d) of figure 2) three different thermal conditions were applied.

- Step 1  (Burning step): On the disc, the temperature was increased to $T_{\text{max}}$ in order to emulate the burning process.

- Step 2  (Reaction time): This step corresponds to the reaction time vis-a-vis the burn. It was limited to 5 seconds. The same boundary conditions were imposed as in step 1, but the burning process was replaced by convection with air. Thus, all the upper part of the dermis was subject to convection.

- Step 3  (Health care): Two approaches were proposed to reduce the damage caused by the burn in order to limit its spread:

+ Treatment by water: The burned zone was under a flow of water during 35 s. The convection was induced by fluid movements and modeled by a forced convection. The convection coefficient can take values between 100 and 15,000 [W.m$^{-2}$.K$^{-1}$]. In our computations the convection coefficient $h_{\text{water}}$ was chosen at a value equal to 500
[W.m⁻².K⁻¹]. It corresponds to a convection due to a flow coming from ordinary tap water of diameter 15 mm and assuming a flux of 45 l/min (ejection velocity of about 4 m/s). For a horizontal laminar plane flow, we used the following expression for the Nusselt number:

\[ N_u = 0.66(Pr)^{1/3}(Re)^{1/2} \]  \hspace{1cm} (5)

where \( Pr \) is the Prandtl number and \( Re \) the Reynolds number. The Nusselt number is linked with \( h_{\text{water}} \) by the following relationship:

\[ h_{\text{water}} = k_f N_u / L \]  \hspace{1cm} (6)

where \( k_f \) is the fluid thermal conductivity (0.6 W/m.K) and \( L \) is the length of the model (10 cm). For a water with a specific heat of 4180 J/kg.K, a mass density of 1000 kg/m³ and a dynamic viscosity of 10⁻³ Pa.s, the Nusselt number is equal to 83. The water temperature was chosen at 10°C, which is an average value coming from ordinary tap water.

+ Treatment by ice: we proposed to simulate an ice apposition on the burned zone. The temperature was maintained equal to \( T_{\text{min}} = 1^\circ \text{C} \) on the burned zone during 35 s. These two treatments were compared with a

+ Treatment by air: Like in step 2 external surface is under air convection during 35 s. It means that no treatment is applied to cure the burn.

Obviously, the modeling we suggested to simulate the action of the burner and the ice (maintaining a constant temperature \( T_{\text{max}} \) or \( T_{\text{min}} \)) is quite oversimple and would require to be refined.

3 Numerical simulations

Concerning the numerical resolution of heat equations (1) and (3), the finite element method was chosen and an implicit time integration scheme was adopted. The mesh contains 13,593 quadratic tetrahedral elements (see figure 3).

This algorithm was programmed using COMSOL and MATLAB softwares. Calculations were carried out with a time step \( \Delta t = t_{i+1} - t_i = 0.25 \) s. The generations of metabolic internal power in the dermis and the hypodermis were fixed at 368 W.m⁻³ (see Roetzel and Xuan, 1998)). The numerical model was validated and was able to estimate blood velocities (see Ratovoson et al., 2011)). The thermal response of the skin depends on its thickness. During the phase of validation of the model, we modified the thickness of the layer of dermis in varying the depth of the vein, because the vein was supposed to be the separation between dermis and hypodermis. We varied the radius of the vein too. The depth of the vein was locally measured with echo Doppler and an average of 1 mm was chosen to best-fit thermal measured responses.

New developments concern the coupling with the thermal damage. The rates of thermal damage integration were computed with an Euler implicit scheme as follows

\[ \Omega(t_{i+1}) = \Omega(t_i) + \Delta t A \frac{\delta p}{RT(t_{i+1})} \]  \hspace{1cm} (7)
Temperatures at time $t_{i+1}$ were given by the previous thermomechanical calculations. The coupling only was considered from the thermomechanical model to the thermal damage model. The influence of thermal damage on thermomechanical properties was not taken into account. We are aware that such simplified assumption is wrong, but currently we are not able to model this coupling.

Two cutting planes of symmetry was defined to show numerical results for thermal damage: the longitudinal plane A-A and the transversal plane B-B (see figure 4).

3.1 Hyperthermia injury evaluation: no health care

Following results were obtained after burning simulations. For all computations, the activation energy of the skin was $E_a = 6.27 \times 10^8$ [J.kmol$^{-1}$]. Figure 5 represents the damage in the longitudinal cutting plane A-A at the end of the simulation. These results were carried out with a factor of frequency $A = 3.1 \times 10^{90}$ s$^{-1}$. Calculations simulated a treatment by air. The result on the left was obtained after applying a burning temperature $T_{\text{max}} = 400^\circ\text{C}$ during an exposure duration $t_\Omega = 10$s. The result on the right was found after applying a burning temperature $T_{\text{max}} = 100^\circ\text{C}$ during an exposure duration $t_\Omega = 5$s. These results are valid for the two most extreme cases of burning. It is noted that for a severe burn, the damage reached the vein and was transported with blood flow.

Left image of figure 6 represents the damage on the upper surface of the model at the end of the simulation, for the severe burn. A light transport of the damage caused by the vein, can be seen at the bottom of the disc. Right image of figure 6 represents the damage
Figure 4: Cutting planes: longitudinal A-A and transversal B-B.

Figure 5: Thermal damage on the longitudinal cutting plane A-A for a burning temperature $T_{\text{max}} = 400^\circ C$ during an exposure time $t_\Omega = 10s$ (left image) and a burning temperature $T_{\text{max}} = 100^\circ C$ during an exposure duration $t_\Omega = 5s$ (right image).

For a severe burn on the transversal cutting plane B-B, the passage of the vein appears very clearly and separates the damage in two lobes. Tissues under the vein are protected.

To study the evolution of thermal damage in the depth of the tissue, we used the check line plotted in the longitudinal cutting plane A-A (see left image of figure 7). In the right image of figure 7 are plotted thermal damage versus time for points located on the check line.
Figure 6: Thermal damage on the upper surface (left image) and on the transversal cutting plane B-B (right image) for a burning temperature $T_{\text{max}} = 400^\circ C$ during an exposure duration $t_\Omega = 10s$

Figure 7: Left image: thermal damage in the longitudinal cutting plane A-A at the end of the simulation and check line; Right image: thermal damage evolution for points located on the check line

The calculations were done with the following values: $A = 3.1 \times 10^{70} \text{ s}^{-1}$, $t_\Omega = 10s$ and $T_{\text{max}} = 400^\circ C$. Several phenomena are observed:

- The maximum damage is located on the surface in contact with the burner (the upper surface).
- The value of this damage exceeds $10^4$, a value which corresponds to the 3rd degree of burn.
- This damage stops brutally and seems not to evolve after 10s. In fact, there is a variation which is not perceptible at this scale ($\Omega \simeq 10^{22}$).
- Indeed, a zoom on curves of figure 7, reported in figure 8, confirms that the tissue continues to be damaged after the end of the burn ($t \geq 10s$).
3.2 Hyperthermia injury evaluation: health care tests

We now propose to study the value of the thermal damage according to the treatments we suggested previously. The factor of frequency $A$, the imposed temperature $T_{\text{max}}$ as well as the burning exposure duration $t_{\Omega}$ play a significant role to identify the damage generated by a burn. So we not only tried to evaluate the evolution of the damage according to these three parameters, but also to compare them in order to propose the best treatment and to prevent the damage from spreading.

Figure 9 represents the various curves of thermal damage through the skin depth at the end of the treatment. In this numerical test, we used a factor of frequency $A = 3.1 \times 10^9$ s$^{-1}$ during a burning exposure duration $t_{\Omega} = 10$s, for various temperatures of burn: 100°C, 250°C and 400°C.

For a maximal burning temperature $T_{\text{max}} = 100^\circ$C, the value of the final depth of damaged tissue was identical for the three treatments. The damaged zone reached 1 mm of depth. On the other hand, for $T_{\text{max}} = 250^\circ$C, the treatment by air remained inappropriate and the treatment by ice seems to be the most effective. Indeed, with this latter alternative, the depth and the intensity of the damage are the smallest. For $T_{\text{max}} = 400^\circ$C, the evolution of the damage was almost identical. Depth of damaged tissue reached approximately 3.25 mm. The skin was almost completely injured.

Figure 10 represents the various curves of thermal damage through skin depth during a hyperthermic lapse of time half equivalent to the preceding test, $t_{\Omega} = 5$s.

With an exposure duration $t_{\Omega} = 5$s and a very high burning temperature $T_{\text{max}} = 400^\circ$C, the damage reached 3 mm depth. For $T_{\text{max}} = 250^\circ$C, the maximum depth of damaged skin was 2.2 mm. For $T_{\text{max}} = 100^\circ$C, one observes that the extent of the damage does not depend on the treatments suggested and reached a depth of 0.5 mm. Here again, the treatment by ice seems to be the most suitable.

Figure 11 represents the various curves of thermal damage through skin depth with a factor of frequency $A = 3.1 \times 10^7$ s$^{-1}$ and a burning exposure time $t_{\Omega} = 10$s.

For a burning temperature $T_{\text{max}} = 100^\circ$C, the damage was null. For $T_{\text{max}} = 250^\circ$C, the
Figure 9: Hyperthermal damage through skin depth (mm) for various temperatures of burn, during an exposure duration $t_\Omega = 10s$ according to health care treatments (air, water, ice) with a factor of frequency $A = 3.1 \times 10^{90} \text{ s}^{-1}$.

Figure 10: Hyperthermal damage through skin depth (mm) for various temperatures of burn, during an exposure duration $t_\Omega = 5s$ according to health care treatments (air, water, ice) with a factor of frequency $A = 3.1 \times 10^{90} \text{ s}^{-1}$.

damage was identical for the three treatments and reached a depth of 1 mm. Finally, for $T_{\text{max}} = 400^\circ \text{C}$, the maximum depth of damaged skin was of 1.65 mm. Numerical results show that with this choice of factor of frequency, the health care treatment has no influence.
Figure 11: Hyperthermal damage through skin depth (mm) for various temperatures of burn, during an exposure duration $t_\Omega = 10\,s$ according to treatments (air, water, ice) with a factor of frequency $A = 3.1 \times 10^{70} \,s^{-1}$

on the depth of damaged skin.

According to the previous results, general comments are the following:

- In the manner of (Becker and Kuznetsov, 2007), we compared the various treatments between them, by changing the conditions of burns (temperature, duration) and the significant parameters of the damage law (activation energy and factor of frequency).
- As reported in (Becker and Kuznetsov, 2005) we showed a transportation of damage along the vein.
- For a high burning temperature $T_{\text{max}} = 400^\circ\text{C}$, and for all values of the factor of frequency $A$ and burning exposure duration $t_\Omega$, the damage was almost identical for the three health care treatments. However, for a short burning exposure duration, the value and the depth of the damaged skin change.
- The burning exposure duration plays a significant role on the extent of the thermal damage on the tissue. For the three results presented above, when burning temperature was not very high ($T_{\text{max}} = 100^\circ\text{C}$), the depth of damaged tissue was identical. The treatments do not affect the extent of the damage.
- Treatments have a significant action only when the burn is fairly important. In this case, the treatment by ice seems to be the most suitable.
- The influence of treatments vanishes when the factor of frequency decrease.
- Contrary to investigations reported in (Kuznetsov, 2006) we did not propose to optimize treatments while varying their parameters (convection coefficient, temperature of water,
temperature of the ice, duration of the treatment). We placed the study under conditions of classical interventions available on a place of accident and we tried to highlight the best treatment between two possible.

- We also showed that the reaction time is even more determining than the treatment for this damage model. Beyond 5 seconds of reaction, it is too late to limit the damage, whatever the applied treatment.

These results are partial. To confirm the tendencies, it would also be necessary to test the influence of the coefficient of transfer $h_{water}$, the temperature of water, the temperature of the ice, the transfer of heat between the ice and the skin, the duration of the reaction time and the duration of the treatment.

4 Conclusion

We developed a numerical model of thermal damage coupled with a 3D multi-layer model including blood perfusion and a vein. A light transport of the damage can be observable. Such a transport is caused by the vein. The passage of the vein appears very clearly. It separates the damaged zone into two lobes and protects tissues that are located under it. In addition, this model of damage allows us to suggest two treatments for the burns, while trying to decrease the spread of the damage. The factor of frequency, the burning temperature, as well as the burning exposure duration play a significant role in identifying the severity of the damage. Calculations showed that the spread of the damage could be decreased. In our examples, it appears that a treatment simulating ice apposition on the burn is more effective than a treatment by flow of water. Our results show that this report is true only under conditions of moderate burn. We also showed that the reaction time is even more determining than the treatment for this damage model. Beyond 5 seconds of reaction, it is too late to limit the damage, whatever the applied treatment. This model opens up new perspectives for future work on an improvement of the treatments of burns.

References


