Analysis of Thermal Relaxation During Laser Irradiation of Tissue

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Background and Objective: Thermal relaxation time ($\tau_r$) is a commonly-used parameter for estimating the time required for heat to conduct away from a directly-heated tissue region. Previous studies have demonstrated that temperature superposition can occur during multiple-pulse irradiation, even if the interpulse time is considerably longer than $\tau_r$. The objectives of this study were (1) to analyze tissue thermal relaxation following laser-induced heating, and (2) to calculate the time required for a laser-induced temperature rise to decrease to near-baseline values.

Study Design/Materials and Methods: One-dimensional (1-D) analytical and numerical and 2-D numerical models were designed and used for calculations of the time $t_{\text{eff}}$ required for the peak temperature ($T_{\text{peak}}$) to decrease to values slightly over baseline ($\Delta T_{\text{base}}$). Temperature values included $T_{\text{peak}} = 65$ and 100°C, and $\Delta T_{\text{base}} = 5$, 10, and 20°C. To generalize the calculations, a wide range of optical and thermal properties was incorporated into the models. Flattop and gaussian spatial beam profiles were also considered.

Results: 2-D model calculations of $t_{\text{eff}}$ demonstrated that $t_{\text{eff}}$ (2-D) was as much as 40 times longer than $\tau_r$. For a given combination of $T_{\text{peak}}$ and $\Delta T_{\text{base}}$, a linear relationship was calculated between $t_{\text{eff}}$ (1-D) and $\tau_r$ and was independent of optical and thermal properties. A comparison of 1-D and 2-D models demonstrated that 1-D models generally predicted longer values of $t_{\text{eff}}$ than those predicted with a 2-D geometry when the laser spot diameter was equal to or less than the optical penetration depth.

Conclusion: Relatively simple calculations can be performed to estimate $t_{\text{eff}}$ for known values of $\tau_r$, $T_{\text{peak}}$ and $\Delta T_{\text{base}}$. The parameter $t_{\text{eff}}$ may be a better estimate than $\tau_r$ of tissue thermal relaxation during multiple-pulse laser irradiation. Lasers Surg. Med. 29:351–359, 2001. © 2001 Wiley-Liss, Inc.

Key words: effective relaxation time; thermal modeling, finite difference; spatial beam profile; heat conduction

INTRODUCTION

Pulsed laser radiation is used to heat tissue and precisely ablate or coagulate tissue. In laser surgery, the general goal is to selectively heat a desired region of tissue while minimizing collateral thermal damage. The theory of selective photothermolysis proposed by Anderson and Parrish [1] is generally applied to a given situation to determine the appropriate laser pulse duration ($t_p$).

From the theoretical analysis of Anderson and Parrish [1], the concept of thermal relaxation time has become a popular term for selection of laser pulse duration. The thermal relaxation time ($\tau_r$) of a heated region of tissue is the time required for the peak temperature rise ($\Delta T_{\text{peak}}$) in a heated region of tissue to decrease to 37% of the total rise:

$$\tau_r = \frac{\delta^2}{4\alpha}$$

where $\delta$ is usually the penetration depth (cm). The appropriate value for $\delta$ is discussed in more detail in the Discussion section. A common belief is that if $\tau_r$ is less than $\tau_p$ of the irradiated region, thermal damage is minimized. However, for procedures in which high peak temperatures ($T_{\text{peak}} > 100^\circ C$) are expected, such as laser skin resurfacing, the temperature value may still be sufficiently large to induce thermal damage after $\tau_r$ has elapsed. For example, assuming an initial temperature of 30°C and a laser-induced temperature rise to 100°C, $T_{\text{peak}}$ will be approximately ($100^\circ C - 30^\circ C$) $0.37 + 30^\circ C = 56^\circ C$ after $\tau_r$ has elapsed. If another laser pulse is applied to the tissue at this point, thermal effects may be more severe due to temperature superposition effects. Complete thermal relaxation of tissue may take several thermal relaxation times [2]. Also, $\tau_r$ is applicable only for specific tissue conditions. Tissue optical and thermal properties may change during the laser pulse and for consecutive pulses due to evaporation and tissue denaturation [3,4]; as these properties change, the value of $\tau_r$ changes as well.

Several theoretical and experimental studies have shown that temperature superposition can occur even if the time between pulses is several orders of magnitude greater than the thermal relaxation time [5–8]. In

Contract grant sponsor: Air Force Office of Scientific Research through MURI from DDR&E; Contract grant number: F49620-98-1-0480; Contract grant sponsor: Texas Higher Education Coordinating Board; Contract grant number: BER-ATP-253; Contract grant sponsor: Albert and Clemmie Caster Foundation.

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Accepted 30 April 2001
previous measurements of skin radiometric surface temperatures during pulsed CO2 laser irradiation (τ_{p} ~700 μs) [5], it was demonstrated that elevated temperatures remained for 20–40 ms after the end of a laser pulse. These results indicate that complete tissue thermal relaxation to baseline temperatures is a relatively slow process.

The purpose of this study was to investigate theoretically the thermal response of soft tissue to laser-induced heating. The time required for T_{peak} to decrease to values just above baseline is calculated using a 2-D numerical thermal model. Results of a 1-D analytical solution to the heat conduction equation are compared to those of the numerical model to demonstrate its applicability as a rule-of-thumb estimation for the effective relaxation time (τ_{rel}) of tissue.

MATERIALS AND METHODS

Conduction heat flow is governed by the heat conduction equation. In three dimensions, conduction heat flux $q$ (W/cm²) is defined as:

$$q = -k \nabla T,$$  \(2\)

where $k$ is thermal conductivity (W/cm/K) and $\nabla$ is the del operator. In Cartesian coordinates,

$$\nabla = \frac{\partial}{\partial x} + \frac{\partial}{\partial y} + \frac{\partial}{\partial z}$$

The negative sign in Equation (2) is due to the direction of heat conduction; heat always flows from a higher temperature region to a lower temperature region.

In cylindrical coordinates, Equation (2) becomes:

$$\frac{1}{r} \frac{\partial}{\partial r} \left( r \frac{\partial T}{\partial r} \right) + \frac{1}{r^2} \frac{\partial^2 T}{\partial \theta^2} + \frac{\partial^2 T}{\partial z^2} + \frac{q_{gen}}{k} = \frac{1}{\alpha} \frac{\partial T}{\partial t}, \quad 3$$

where $q_{gen}$ is the laser source term (W/cm³) and $\alpha$ is thermal diffusivity (cm²/s). If 1-D heat flow is assumed, Equation (2) takes on the following form:

$$\frac{\partial^2 T}{\partial z^2} + \frac{q_{gen}}{k} = \frac{1}{\alpha} \frac{\partial T}{\partial t}. \quad 4$$

2-D Finite Difference Model

A finite difference solution [9] to the heat conduction equation (Equation (2)) was derived. Cylindrical coordinates were used, resulting in a 2-D axisymmetric tissue geometry (Equation (3)). The tissue was assumed to be homogeneous and its dimensions were 1-cm thickness and 1-cm radial extent. The size of each grid element was 40 × 20 μm.

Accurate thermal modeling of tissue subjected to relatively high temperatures is difficult due to potential dynamic changes in optical and thermal properties as a function of temperature and water content [3,4]. For the 2-D simulations, we assumed that tissue consisted of 70% water (representative of dermis [10]) and calculated thermal properties from the following equations [11]:

$$\rho = (6.16 \times 10^{-2} W + 0.938)^{-1} \quad 4a$$

$$c = 2.5 W + 1.7 \quad 4b$$

$$k = \rho \times 10^{-2} (0.454 W + 0.174) \quad 4c$$

where $\rho$ is density (g/cm³), $c$ is specific heat (J/g/K), and $W$ is water content (e.g., for 70% water, $W = 0.7$). For the 1-D simulations (described below), other sets of thermal properties were considered that corresponded to different water contents.

Absorption coefficient ($\mu_a$) values of 10, 100, and 1000 cm⁻¹ were selected. These values were representative of those commonly encountered during laser-mediated tissue heating. Intermediate absorption coefficients were used in the 1-D simulations (described below). Three different spot diameters ($a_o = 500 \mu$m, 1.5 mm, and 2.5 mm diameters) were used to take into account various spot size/penetration depth ($\delta = 1/\mu_a$) combinations. Flattop and gaussian spatial beam profiles were considered. For gaussian profiles, $a_o$ represents the 1/e² diameter.

Pulse durations were chosen to correspond to 1/100th of the thermal relaxation time. This rule for pulse duration selection was chosen since it is reasonable to assume that such a value would generally be considered as sufficient for limiting heat conduction during the pulse.

Radiant exposures were chosen so that the peak temperature $T_{peak}$ at the end of the laser pulse corresponded to either 65 or 100°C. The value of 65°C was chosen because it corresponds to a commonly-cited threshold value for collagen denaturation [7,12,13], although in reality thermal damage is a temperature-time phenomenon [14]. 100°C corresponds to water boiling at atmospheric pressure and is a good estimate of the residual peak temperature in tissue immediately after ablation. The effective relaxation time ($\tau_{rel}$) required for the peak temperature to fall to $\Delta T_{base} = 10^0$C above baseline ($T_{baseline} = 22^0$C) was calculated for each model run.

1-D Analytical Expression

An advantage of using 2-D models is that they offer more realism than 1-D models. A major disadvantage is the considerably longer computation time required to arrive at a solution. A 1-D equation derived by Anvari et al. [15] from basic principles presented by Carslaw and Jaeger [16] describes 1-D heat flow in a semi-infinite medium with an adiabatic surface boundary condition. This equation was slightly modified to the following form:

$$\Delta T(z,t) = \frac{H_o \mu_a}{2 \rho c} \exp(-\xi^2) [\text{erfc}(\beta - \xi) + \text{erfc}(\beta + \xi)], \quad 5$$

where $z$ is depth (cm), $t$ is time (s), $H_o$ is the incident radiant exposure (J/cm²), $\xi = z/2 \sqrt{\tau \beta}$, $\beta = \mu_a \sqrt{\tau}$. $1/2$ erfc is the complementary error function, and $\text{erfc}(x) = \exp(x^2) \text{erfc}(x)$. Equation (5) is valid for an initial temperature distribution that is proportional to $\exp(-\mu_a z)$; in other words, heat conduction during the pulse is assumed to be negligible. Equation (5) was first compared to results
of a 1-D implicit finite difference model in which heat conduction during the pulse was assumed to determine the relative accuracy of this equation. Results of this equation were then compared to those obtained from the 2-D model to determine its suitability for predicting $t_{eff}$.

Since the time required to implement Equation (5) was considerably shorter than that required with the 2-D finite difference equations, this equation was applied to a wider variety of cases. Tissue thermal properties were evaluated for four different tissue scenarios: (1) 100% water, (2) 70% water, (3) 0% water, and (4) pure carbon. These sets of thermal properties represented asymptotic limits of tissue states encountered during high-power laser irradiation of tissue and are compiled in Table 1. Equations (4a)–(4c) were used to calculate thermal properties for the first three cases; pure carbon thermal properties were taken from various sources [9,17,18].

The following absorption coefficients were used: 10, 20, 30, 40, 50, 100, 200, 300, 400, 500, 600, 700, 800, 900, and 1000 cm$^{-1}$. Radiant exposures were chosen so that peak temperatures of 65 and 100°C were obtained by using the following equation:

$$H_o = \frac{\Delta T_{peak} \rho c}{\mu_a},$$

(6)

where $\Delta T_{peak} = T_{peak} - T_{baseline}$ is the peak temperature rise (°C). Equation (6) is simply a 1-D solution to the heat conduction equation at the surface and in the absence of conduction.

The time required for the peak temperature to decrease to $T_{base}$ values of either 5, 10, or 20°C above baseline were calculated. The calculations for 10°C above baseline were directly comparable to the 2-D model runs.

RESULTS

2-D Finite-Difference Model

A total of 36 simulations were run. Figure 1 depicts surface temperature profiles for a representative model run. In the 2-D simulations, $\alpha = 1.42 \times 10^{-2}$ cm$^2$/s. $H_o$ was

![Fig. 1. Surface temperature profiles for a model run at $t = 17.6$ ms (solid line), which is the end of the incident laser pulse, and $t = 9.567$ seconds (dashed line), which is the point in time at which the peak temperature is 32°C. $H_o = 8.16$ J/cm², gaussian beam profile, spot radius = 2.5 mm, $\mu_a = 10$ cm$^{-1}$.](image-url)
chosen so $\Delta T_{\text{peak}}$ at the end of a 17.6-ms pulse was 65°C. The time required for $T_{\text{peak}}$ to decrease to 32°C was 9.57 seconds. For the experimental conditions simulated in this case, $\tau_r$ was 1.76 seconds. Thus, in this example, $\tau_{\text{eff}}$ was 5.44 times longer than $\tau_r$.

A summary of model results is provided in Table 2. The dimensionless variables $\Gamma$ and $\tau$ represent the ratios $\omega_0/\delta$ and $\tau_{\text{eff}}/\tau_r$, respectively. Note that for all but one model run, $\tau$ is greater than unity, indicating that the times required for $T_{\text{peak}}$ to decrease to 32°C were longer than $\tau_r$.

Figure 2 shows a plot of $\tau_{\text{eff}}$ calculated for gaussian beam profiles as a function of $\tau_{\text{eff}}$ for flattop beam profiles. Each coordinate pair represents identical simulated experimental conditions except for the spatial beam profile (Table 2). The solid line represents the boundary at which $\tau_{\text{eff},\text{gaussian}} = \tau_{\text{eff},\text{flattop}}$. All data points fall below this line, indicating that for a given set of experimental conditions, the value of $\tau_{\text{eff}}$ required for gaussian temperature distributions to thermally relax is shorter than that for uniform distributions. The discrepancy is larger for longer values of $\tau_{\text{eff}}$.

### 1-D Analytical Equation

Equation (5) was applied to a total of 456 different experimental scenarios. Calculations of $\tau_{\text{eff}}$ for relatively short pulses ($\tau_p = 0.01 \tau_r$) were compared to those obtained from 1-D implicit finite difference calculations and agreed to within 1%. Results of all model runs are shown in Figure 3. Note that all calculations of $\tau_{\text{eff}}$ fall above the solid line, indicating that the times required for $T_{\text{peak}}$
decreasing to values 5–20°C above baseline are longer than \( t_r \).

In Figure 3, it is evident that for given combinations of peak temperature rise (\( T_{\text{peak}} = 43 \) or 78°C) and final peak temperature above baseline (\( \Delta T_{\text{base}} = 5, 10 \) or 20°C), a plot of \( t_{\text{eff}} \) vs. \( t_r \) yields data that fall along straight lines. Average slopes of linear fits are listed in Table 3. The slope of each line dictates the offset of each line on the plot; a larger slope indicates a higher offset. Note that for a given combination of \( T_{\text{peak}} \) and \( T_{\text{base}} \), the linear slopes are identical regardless of the combination of tissue optical and thermal properties used in the model.

**Comparison of 2-D and 1-D Model Results**

Figure 4 shows \( t_{\text{eff}} \) (1-D model)/\( t_{\text{eff}} \) (2-D model) as a function of \( t_{\text{eff}} \) (2-D model). In general, the 1-D model overpredicts \( t_{\text{eff}} \) (e.g., y-axis values greater than unity). This overprediction is associated with the ratio (\( \Gamma \)) of spot size (\( \omega_0 \)) to penetration depth (\( \delta \)). When \( \omega_0 \gg \delta \) (e.g., for large \( \Gamma \)), the 1-D solution is accurate at the center of the laser spot, and thus the values of \( t_{\text{eff}} \) calculated with the 1-D and 2-D models are comparable (Fig. 5).

**DISCUSSION**

The main advantage of \( t_r \) is that it is a simple quantity to calculate. It can also be used to adequately predict thermal effects from single-pulse irradiation procedures, such as Port Wine Stain treatment [19]. For cases in which multiple pulses are delivered, such as laser skin resurfacing or laser-assisted blepharoplasties, our results demonstrate that \( t_r \) is not a good indicator for estimating thermal effects (Fig. 3 and Table 2). Model calculations predict that \( t_{\text{eff}} \) (2-D) can be as much as 40 times longer than \( t_r \) for peak temperature relaxation from 100 to 32°C for a 22°C baseline temperature; other researchers have estimated comparable values of \( t_{\text{eff}} \) [7, 8].

Since \( t_{\text{eff}} \) can be on the order of 30 seconds (e.g., model run #6 in Table 2), it is necessary to consider the effects of blood perfusion on \( t_{\text{eff}} \). Welch et al. [20] estimated that perfusion has a significant effect on temperatures in normal skin after approximately 100 seconds of laser heating. Thus, the values of \( t_{\text{eff}} \) calculated in this study are not expected to differ significantly if blood flow were considered.

van Gemert and Welch [21] demonstrated that the value of \( \delta \) in Equation (1) depends on \( \Gamma = \omega_0 / \delta \). For large values of \( \Gamma \), \( \delta \) is the penetration depth of light. For small values of \( \Gamma \), \( \delta \) is better represented by \( \omega_0 \). Many laser-based therapeutic procedures involve conditions in which \( \Gamma \) is large, and so \( \delta \) is the penetration depth. To minimize confusion, in this study we assumed that \( \delta \) is the penetration depth for all values of \( \Gamma \).
Data in Figure 2 suggest that for a given parameter set, thermal relaxation is a faster process for incident gaussian beams than for flattop beams. This can be explained by a discussion on radial temperature gradients. Normalized, representative temperature profiles at the end of laser irradiation are shown in Figure 6 for incident gaussian and flattop beams that are 2.5 mm in radius. Initially, at \( r = 0 \), no net radial heat conduction will occur for the flattop case because the temperatures at adjacent radial positions are equal in magnitude. On the other hand, for the gaussian example, net heat flow away from the central axis occurs almost immediately because adjacent regions (e.g., \( r > 0 \)) are at lower temperatures. The discrepancy between \( t_{\text{eff}} \) for gaussian and flattop beams increases as \( \mu_a \) gets larger because, for a given \( \mu_a \), as spot size increases, the region of \( T = T_{\text{peak}} \) during flattop-beam irradiation increases as well. For a gaussian beam, \( T_{\text{peak}} \) occurs only at the central axis and its location is insensitive to changes in spot size.

Schomacker et al. (7) observed that heat required for ablation during CO\(_2\) laser ablation of in vitro guinea pig skin increased from 4.8 to 6.6 kJ/g as \( \omega_0 \) decreased from 680 to 250 \( \mu \)m. Our model results (Table 2) show that for a given \( \mu_a \), \( t_{\text{eff}} \) (2-D) decreases as \( \omega_0 \) gets smaller, suggesting that heat loss from the irradiated region is faster for smaller beam diameters. This hypothesis offers a reasonable explanation for why a decrease in \( \omega_0 \) leads to an increase in heat required for ablation.

A plot of \( t_{\text{eff}} \) (1-D) calculated using Equation (5) as a function of \( \tau_r \) (Fig. 3) shows that a 1-D thermal model...
predicts \( \tau_{\text{eff}} (1-D) > \tau_r \) for the range of \( \tau_r \) considered in this study. The linear relationship between \( \tau_{\text{eff}} (1-D) \) and \( \tau_r \) suggests that a simple multiplicative factor can be used to calculate \( \tau_{\text{eff}} (1-D) \). The value of this factor depends solely on \( \Delta T_{\text{peak}} \) and \( \Delta T_{\text{base}} \) and was calculated for six combinations of \( \Delta T_{\text{peak}} \) and \( \Delta T_{\text{base}} \) (Table 3). Further derivation of Equation (5) demonstrates the reason for this linear relationship. For all times \( t > 0 \), the peak temperature induced by laser light absorption occurs at the tissue surface because the following conditions were assumed during model runs: (1) light scattering was negligible and (2) convective cooling was not considered. Thus, at \( z = 0 \) and at \( t = \tau_{\text{eff}} \), Equation (5) becomes

\[
\Delta T (0, \tau_{\text{eff}}) = \frac{H_{\text{d} \mu_{\text{a}}}}{\text{pc}} \text{erfcx}(\beta).
\]

(7)

In the models, \( \Delta T(0, \tau_{\text{eff}}) \) is equivalent to \( \Delta T_{\text{base}} \) and the quantity \( H_{\text{d} \mu_{\text{a}}}/\text{pc} \) equivalent to \( \Delta T_{\text{peak}} \). The term \( \text{erfcx}(\beta) \) takes into consideration the dynamic change in peak temperature over time. Thus, Equation (7) takes on the following form

\[
\Delta T_{\text{base}} = \Delta T_{\text{peak}} \text{erfcx}(\beta).
\]

(8)

Using Equation (1) and \( t = \tau_{\text{eff}} \), \( \beta \) can be rewritten as

\[
\beta = \left( \frac{t}{\tau_{\text{eff}}} \right) = \mu_{\text{a}}(x \tau_{\text{eff}})^{1/2} = \frac{1}{2} \left( \frac{\tau_{\text{eff}}}{\tau_r} \right)^{1/2}.
\]

(9)

Thus, \( \beta \) represents the ratio \( \tau = \tau_{\text{eff}}/\tau_r \). Since \( \text{erfcx}(\beta) \) is a monotonically-decreasing function as \( \beta \) increases, for a given set of values for \( \Delta T_{\text{base}} \) and \( \Delta T_{\text{peak}} \), there exists a single value for \( \beta \) that satisfies Equation (8). In this situation, \( \beta \) is a constant, so the ratio \( \tau_{\text{eff}}/\tau_r \) is also constant. This relationship is true for any combination of optical and thermal properties. Table 3 provides examples of the linear relationship existing between \( \tau_{\text{eff}} \) and \( \tau_r \) for six combinations of \( \Delta T_{\text{base}} \) and \( \Delta T_{\text{peak}} \).

A comparison between \( \tau_{\text{eff}} \) values calculated from the 1-D and 2-D models (Fig. 4) indicates that the 1-D model is comparable to or overestimates \( \tau_{\text{eff}} \) (2-D). The amount of overestimation depends on \( \Gamma \) (Fig. 5). As described above, for larger spot sizes, radial heat conduction is not as significant as for smaller spot sizes. For \( \Gamma > 10 \), \( \tau_{\text{eff}} \) is equivalent for 1-D and 2-D model calculations; this condition is generally satisfied for incisional or ablation procedures. These results show that the 1-D model can at least provide a conservative estimate of the time required between pulses to minimize potentially deleterious effects of temperature superposition. A worst-case value for \( \tau_{\text{eff}} \) can be calculated using a wide range of thermal properties (Equations 4 (a–c), Table 1). Equation (5) is more complex than the equation for \( \tau_r \) (Equation (1)), but it can still be solved relatively easily using conventional spreadsheet programs.

Since \( \tau_{\text{eff}} \) is linearly related to \( \tau_r \), the accuracy of both values suffers from the same limitation that accurate knowledge of local optical and thermal properties is
required. In particular, to calculate \( \beta \), values of \( \mu_a \) and \( \mu_s \) must be known. Since \( \tau_{\text{eff}} \) is longer than \( \tau_r \), the relative error associated with the uncertainty in a given set of optical and thermal properties is smaller for calculation of \( \tau_{\text{eff}} \) than for \( \tau_r \). It is necessary to keep in mind that both \( \tau_{\text{eff}} \) and \( \tau_r \) are guidelines. The primary purpose of \( \tau_{\text{eff}} \) is to provide an estimate of the time required for heated tissue to cool from \( T_{\text{peak}} \) to a value \( \Delta T_{\text{base}} \) above baseline, whereas \( \tau_r \) represents the time required to cool from \( T_{\text{peak}} \) to a 1/e value.

![Fig. 5. Semilog plot of ratio of spot diameter and penetration depth vs. \( \tau_{\text{eff}} \) (2-D).](image)

![Fig. 6. Normalized temperature profiles at the end of a laser pulse for flattop (solid line) and gaussian (dashed line) beam profiles.](image)
CONCLUSIONS

Theoretical calculations showed that the concept of thermal relaxation time does not adequately provide a measure of time that can be used during multiple-pulse irradiation of tissue. It is necessary to consider the slow temperature decay that occurs for times longer than $t_r$. The following equation can be used to estimate $t_{\text{eff}}$.

$$t_{\text{eff}} = m t_r = \frac{m \delta^2}{4\nu},$$

where $m$ is the slope of the line fitting $t_{\text{eff}}$ (calculated with Equation (5)) as a function of $t_r$. Calculated values of $m$ for six combinations of $\Delta T_{\text{peak}}$ and $\Delta T_{\text{base}}$ are provided in Table 3; it is straightforward to calculate $m$ with Equation (5) for other combinations.

REFERENCES