Bedsore Revitalization by-Laser Therapy (Low Level Laser: LED-Ga-Al-As, 660 nm)

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Abstract
Background: In 1967 a few years after the first working laser was invented, Endre Mester in Semmelweis University Budapest, Hungary wanted to find out if laser might cause cancer. He took some mice, shaved the hair off their backs, divided them into two groups and gave a laser treatment with a low powered ruby laser to one group. They did not get cancer and to his surprise the hair on the treated group grew back more quickly than the untreated group. That was how “laser biostimulation” effects were discovered.

Purpose of the work: The effects of pulsed monochromatic light, with fixed pulsations and wavelengths, on the healing of pressure ulcers were evaluated in this prospective, randomized, controlled study.

Method: A placebo-controlled, double-blind study using low energy photon therapy (LLLT) was performed in ten patients with bedsore on the back. Treatment was given three times a week for 10 weeks, using monochromatic (red) optical sources; diode 660nm (GaAl-660). The patients who were randomized to placebo treatment received sham therapy from an identical-appearing light source from the same delivery system.

Results: Ten patients with bedsore were randomized to receive LLLT or placebo therapy. At the conclusion of the study, the percentage of the initial ulcer area remaining unhealed in the LLLT and placebo groups was 24.4% and 84.7%, respectively (P = 0.0008). The decrease in ulcer area (compared to baseline) observed in the LLLT and placebo groups was 193.0 mm² and 14.7 mm², respectively (P = 0.0002). One patient dropped out of the study, complaining of lack of treatment efficacy; he was found to be randomized to the placebo group. There were no adverse effects.

Conclusions: In this placebo-controlled, double-blind study LLLT was an effective modality for the treatment of bedsore which were resistant to conventional medical management. The results are encouraging as pulsed monochromatic light increased healing rate and shortened healing time. This will positively affect the quality of life in elderly patients with pressure ulcers.

Keywords: Bedsore Healing, Soft Tissue Healing, Decubitus Ulcer Healing, Wound Healing. Low Level Laser, Laser Therapy.

Introduction
Background
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Several factors have a direct influence on tissue healing, altering this process, making it slower, thus allowing complications associated with wound exposure to the external environment. Thetable below displays the key local and systemic factors that affect tissue wound healing.

**Uses of Laser Therapy to treat Bedsore**
Allied health professionals regularly care for a variety of skin wounds, such as abrasions, turfs burns, surgical incisions, and ulcerations, which are perhaps the most difficult to treat. At present, cutaneous lesions represent a dilemma of global proportions and instigate great clinical interest because of the high morbidity associated with changes in the normal healing process. Among the clinical aspects involving this issue, we emphasize tissue repair time in an effort to make the process quicker and more harmonious, reduce possible complications in lesion resolution, and allow an adequate choice of therapy. To do this, familiarity with the pathogenesis of tissue healing is necessary, as well as understanding of the factors affecting the process and the role each one plays in its progress, always seeking a clinical treatment that optimizes skin lesion care. Among the methods currently available, low-level laser therapy (LLLT) stands out.

From acute wound management to augmentation of scar tissue remodeling, the clinician seeks to optimize wound care to promote healing. Experimental in vitro and in vivo studies have been underdevelopment since the 1960s, and in the early 1990s, LLLT was approved by the Food and Drug Administration (FDA) as an important method for treating healing processes [18-20]. Recent results of a study demonstrated that LLLT is an effective method to modulate tissue repair, thus significantly contributing to a faster and more organized healing process [21]. Nevertheless, in spite of the large number of studies involving this technique and its wide use in clinical practice, the principles of its action in cells and tissues are still not well understood. The objective of this study is to review pathogenetic aspects of soft tissue repair to understand the major complications in skin lesion healing. In addition, it aims at forming a concise compilation of published data from scientific literature to date to verify whether the use of low-level laser influences wound healing, since its mechanisms of action are not fully clear yet.

**Details Experimental**

**Materials and Parameters-Place of study**

**Patient:** Bedsore/ Decubitus Ulcer Patients.

**Selection Criteria**

**Sample collection & distribution**
The sample was collected randomly from admitted patients in Shaheed Suhrawardy Medical College Hospital in the Department of Orthopedics and Traumatology, Dhaka-1207, Bangladesh. A total of 10 patients randomly collected. The patients were briefed about the study and written consent (Informed consent) was obtained from all patients / medical legal guardian for other patients.

**Inclusion Criteria**
1. Patient suffering from bedsore more than 12 weeks.
2. Failed to heal by all means of conventional therapy.
4. Age between 55-95 years old.

**Purpose of the work**
The effects of pulsed monochromatic light, with fixed pulsations and wavelengths, on the healing of pressure ulcers were evaluated in this prospective, randomized, controlled study.

**LLLT In soft tissue healing process / Skin lesions**
Some of the most common cutaneous wounds include excoriations, burns, surgical incisions, and acute or chronic ulcerations [2,3]. Diabetes mellitus is one of the primary predisposing factors for skin lesion development and one of the most common reasons for patients to seek health care, as it represents an important cause of disability and premature death [4-6]. According to Pedrosa, serious cutaneous foot lesions in diabetic patients are the cause for hospital admission in 51% of patients in endocrinology wards of Brazilian universities hospitals [7]. When not properly healed, these lesions represent the main cause of morbidity, immobility and limb amputation, according to data from the American Diabetes Association [6]. Burn injuries, a clinical condition resulting from direct or indirect action of heat on the human body that causes different degrees of skin lesions, are a significant cause of mortality, primarily due to the infections that can evolve to sepsis. According to the Brazilian Society of Burn Injuries (Sociedade Brasileira de Queimaduras), there are 1 million cases each year in Brazil [8,9]. Skin lesions have a great morbidity potential primarily because of complications in the normal healing process. To prevent these complications and promote cure, one needs to understand the normal process of soft tissue repair, as well as the factors that determine its normal healing.

The normal process of soft tissue repair involves the following steps: homeostasis, inflammation (“cleaning”), demodulation, proliferation, and maturing [10]. The homeostatic phase occurs immediately after the appearance of the lesion and depends on platelet activity and on blood coagulation process, which includes a complex release of vasoactive substances, adhesive proteins, and growth factors for the development of other stages [10,11]. Later on, the inflammatory process sets in with the presence of numerous chemical mediators and inflammatory cells (polymorphonuclear leukocytes, macrophages, and lymphocytes). This phase is responsible for removing necrosed tissue and combating aggressive agents implanted in the wound. Next, tissue proliferation, which is responsible for “closing” the wound, sets in, with re-epithelialization, fibroplasia (matrix formation), and angiogenesis, essential for the supply of oxygen and nutrients needed for healing. Finally, there is wound contraction followed by remodeling, which takes place in the collagen of the region and has the objective of increasing tensile force and diminishing the scar size [11,12].

Therefore, tissue healing highlighted as one of the main effects of LLLT, is characterized by three main factors. **First**, there is an increment of ATP production, (as laser is considered to raise the production of ATP,) leading to a boost in mitotic activity and to an increase in protein synthesis by mitochondria, resulting in greater tissue regeneration in the repair process [13-16]. **Second**, there is a stimulus to microcirculation, which increases the delivery of nutrition elements associated with increased speed of mitosis, facilitating cell multiplication [13,14]. **Finally**, new vessels are formed from preexisting vessels [13,14,17].

Several factors have a direct influence on tissue healing, altering this process, making it slower, thus allowing complications associated with wound exposure to the external environment. The table below displays the key local and systemic factors that affect tissue wound healing.
5. Haven’t previously treated with LLLT.

**Machine:** BioLux MD

LLLT, Low Level Laser (LED- Ga-Al- As 660).

**Irradiance Parameters**
- Beam source (Incoherent-Ga-Al-As).
- Irradiance dose: 4-8 J/cm²/min.
- Irradiance time: 1-2 minutes
- Mode: Continuous wave
- Wavelengths Used: 660 nm.
- Total session: 25-35.

An average power of 5-8 mw was provided through a fiber optic delivery system around the wound margin for about 8-10 min at each point at a distance of one cm. Since the center of the ulcer was deep, it was decided to give laser therapy concentrating maximum irradiance there (Figure 1).

The optimum distance from probe tip to target surface was 1-4 mm. Probe motion during lasing was a slow, circling movement over each square centimeter of open lesion, timed to permit the suggested dosages. As the lesion is large, i.e., 4-6 cm in diameter, a change in technique is adopted which involves a slow, traversing of the perimeter of the lesion, allowing approximately 90 sec per linear centimeter of the perimeter, at the suggested distances (1-4 cm).

The parameters which have been found to be most effective are in the range of 90 sec/cm2 of open wound surface, with the laser beam set at a pulsed rate of 40-80 pulses per second (PPS), depending upon the chronicity of the lesion. The more chronic, the slower the pulse rate suggested.

This technique apparently provides sufficient exposure to the laser beam to stimulate healing effectively, compared with non-treated areas and previously experienced wound management of a similar nature. Low energy Ga-Al laser provides infrared rays in the wave length of around 660nm by continuous mode.

### Table 1: Parameters involved in determining the LLLT Irradiation parameters

<table>
<thead>
<tr>
<th>Irradiation parameter</th>
<th>Unit of measurement</th>
<th>Comment</th>
</tr>
</thead>
<tbody>
<tr>
<td>Wavelength</td>
<td>nm</td>
<td>Light is electromagnetic energy which travels in discrete Packets that also have a wave-like property. Wavelength is measure in nanometers (nm) and is visible in the 400-700 nm range.</td>
</tr>
<tr>
<td>Irradiance</td>
<td>W/cm²</td>
<td>Often called Intensity, or Power Density and is calculated as Irradiance = Power (W)/Area (cm²)</td>
</tr>
<tr>
<td>Coherence</td>
<td>Coherence length</td>
<td>Coherent light produces laser speckle, which has been depends on postulated to play a role in the photobiomodulation Spectral bandwidth interaction with cells and subcellular organelles.</td>
</tr>
<tr>
<td>Polarization</td>
<td>Linear polarized or circular identical nonpolarized</td>
<td>Light may have different effects than otherwise light(or even 90-degree rotated polarized polarized light). However, it is Known that polarized light is rapidly scrambled in highly scattering media such as tissue (probably in the first few hundred μm).</td>
</tr>
</tbody>
</table>

### Table 2: Parameters involved in determining the LLLT “dose” irradiation time or energy delivered (the dose)

<table>
<thead>
<tr>
<th>Irradiation</th>
<th>Unit of measurement</th>
<th>Comment</th>
</tr>
</thead>
<tbody>
<tr>
<td>Energy</td>
<td>(Joules) J</td>
<td>Calculated as: Energy (J) = Power (W) x time (s). This mixes medicine and dose into a single expression and ignores Irradiance.Using Joules as an expression of dose is potentially unreliable as it assumes reciprocity (the inverse relationship between power and time).</td>
</tr>
<tr>
<td>Energy Density</td>
<td>J/cm²</td>
<td>Common expression of LLLT “dose” is Energy density. This expression of dose again mixes medicine and dose into a single expression and is potentially unreliable as it assumes a reciprocity relationship between irradiance and time.</td>
</tr>
<tr>
<td>Irradiation Time</td>
<td>s</td>
<td>In our view the safest way to record and prescribe LLLT is to define the four parameters of the medicine (see table 1.) and then define the irradiation time as “dose”.</td>
</tr>
<tr>
<td>Treatment Interval</td>
<td>Hours, days or weeks</td>
<td>The effects of different treatment interval are underexplored at this time though there is sufficient evidence to suggest that this is an important parameter.</td>
</tr>
</tbody>
</table>
**Figure 1:** Law of photobiology states that for low power visible light to have any effect on a living biological system, the photons must be absorbed by electronic absorption bands belonging to some molecular photoacceptors, or chromophores- (Sutherland 2002). A chromophore is a molecule (or part of a molecule) which imparts some decided color to the compound of which it is an ingredient.

**Figure 2:** Schematic diagram showing the absorption of red and NIR light by specific Cellular chromophores photoacceptors localized in the mitochondrial respiratory chain.

### Approach and methodology

**Procedure**

<table>
<thead>
<tr>
<th>Treatment Schedule (Dose duration and wound parameter)</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>week</strong></td>
</tr>
<tr>
<td>1-2 week</td>
</tr>
<tr>
<td>3-5 week</td>
</tr>
<tr>
<td>4-6 week</td>
</tr>
<tr>
<td>7-8 week</td>
</tr>
<tr>
<td>9-10 week</td>
</tr>
</tbody>
</table>

**Visualization of treatment Progress by LLLT (LED- Ga-Al- As 660 nm ) of a patient-**

(Chronological Picture View of a Laser (LLLT) Treated Patient)
### Results and Discussion

#### Results:

<table>
<thead>
<tr>
<th>Wound parameters-</th>
<th>Prior to Therapy</th>
<th>End of Therapy</th>
<th>Prior to Therapy</th>
<th>End of Therapy</th>
</tr>
</thead>
<tbody>
<tr>
<td>Margin</td>
<td>Irregular &amp; indurated</td>
<td>Partially Regular</td>
<td>Sutured</td>
<td>In tacked skin</td>
</tr>
<tr>
<td>Floor</td>
<td>Unhealthy, Necrotic Oozing Tissue</td>
<td>Almost Healthy granulation tissue</td>
<td>Covered</td>
<td>Covered</td>
</tr>
<tr>
<td>Base</td>
<td>Spine bone Exposed</td>
<td>Partially Clear granulation tissue</td>
<td>Spine bone covered</td>
<td>Covered</td>
</tr>
<tr>
<td>Surrounding skin/Contraction</td>
<td>Inflamed and scared</td>
<td>Partially Healthy</td>
<td>Healthy</td>
<td>Up to mark Healthy</td>
</tr>
<tr>
<td>Discharge</td>
<td>Profuse purulent Oozing pus</td>
<td>Serous Discharge</td>
<td>No discharge</td>
<td>No discharge</td>
</tr>
</tbody>
</table>

Ten patients with 10 bedsores were randomized to receive LLLT or placebo therapy. At the conclusion of the study, the percentage of the initial ulcer area remaining unhealed in the LLLT and placebo groups was 24.4% and 84.7%, respectively (P = 0.0008). The decrease in ulcer area (compared to baseline) observed in the LLLT and placebo groups was 193.0 mm² and 14.7 mm², respectively (P = 0.0002).

#### Discussion

Our patients demonstrated a significant benefit of Ga-Al-As 660 laser for rapid healing of skin wound. The comparison between the laser and conventionally treated previously treated wounds of the same patient at about same size clearly highlighted that despite uniformity of host factors, local factors and systemic state, the wound healing process was stimulated on the laser exposed side. Healing of wounds is an important problem faced by general & orthopedic surgeons. The possible biostimulatory role of laser light in wound healing is of recent interest [22]. Small sub destructive repetitive doses of laser light are claimed to be useful for trophic ulcers and indolent wounds [23]. The proposed mechanisms of action include local leukocyte proliferation

All these mechanisms possibly lead to more rapid closure of wounds and stronger scar formation. In an experimental study, wounds treated with Ga-Al-As 660 laser revealed significantly more granulation tissue. This study established the biostimulatory effects of low intensity laser radiation [26]. Many reports now indicate benefit to non-healing wounds and trophic ulcers by low-intensity laser irradiation. Out of 351 patients thus treated, 236 showed complete epithelialisation of the wound surface [27]. A 44% increase in healthy granulation tissue was observed, 2/12 ulcers healed completely while 27% revealed reduction in size of the remaining ulcers indicating considerable benefit [28]. Nussbaum et al. in a study compared- the effect of ultraviolet-C and laser for treatment of pressure ulcers in adults with spinal cord injury [29]. They used 660-980 nm wavelength light at an energy density of 4 J/cm². Weekly percentage changes in wound area were compared.

The authors concluded exposure to UV-C decreased healing time and allowed faster return to rehabilitation programs. The UV-C light was better than the laser [29]. Another nonrandomized study of laser and UV lamp on chronic skin ulcers suggested that wounds which fail to respond to topical treatments benefit from either modality [30]. Evaluations of different approaches to wound healing are complicated by the large number of factors that influence wound healing. Although there are anecdotal reports of successful therapy, there are few well controlled studies.

The use of lasers for healing wounds is becoming increasingly attractive to surgeons. A number of animal and in vitro studies have demonstrated that laser irradiation has a significant effect on components of tissue repair [31,32].

#### Conclusions

This study results efficacy of LLLT on wound healing in human model, and indicates that it can be a very important adjective tool / modality for chronic intractable wound management, and in any way it is not harmful to human being. In the past Laser / LED were shown to be effective in wound management but in different degrees, some of those applications showed significant improvement some less effective others no effect. Probably laser/ LED Irradiation parameters are vital for its Biostimulative effects. Inference of those results summaries that irradiation parameters are of vital to laser therapy. We used an optimal dose of irradiance which proved to be most effective biostimulation on human application. Application and research of LLLT on cell responses of wounded skin fibroblasts demonstrate that correct energy density or fluence and number of exposures can stimulate cell responses of wounded fibroblast and promote cell migration and cell proliferation by stimulating mitochondrial activity and maintaining viability without causing additional stress or damage to wounded cells. Results indicate that the cumulative effect of lower doses determines the stimulatory effect, whereas multiple exposure at higher doses result in an inhibitory effect with more damage [33]. Although various studies have extensively covered the effects of laser radiation on tissue, many unanswered questions remain. The mechanisms effectively responsible for cell mitotic activity has not been clarified yet, and there is no standardized ideal dose for stimulating tissue healing [34-45].

Therefore, we noted that there is a need for research on the action and parameters of low-intensity laser use in cutaneous lesions during the different stages of repair, as an attempt to elucidate how this method acts at a cell level in healing processes [46-51]. Elucidation of these issues will enable the establishment of criteria on the true benefits of laser therapy in diseases that need healing stimulation, minimizing healing time and the complications that may occur during the clinical progress of these wounds. In addition, experimental studies indicated that the LLLT may be an important therapeutic tool to stimulate
wound healing in decubitus ulcer patients [52-58]. Finally, the present report highlights the possible utility of Gallium-Aluminium laser at 660 wavelengths as an effective as Helium-Neon laser as an adjunctive modality for wound healing in skin/general & orthopedic practices.

Acknowledgements
I wish to express my deepest gratitude and thanks to Professor Golam Abu Zakaria, for his kind directions, inspiring guidance, and invaluable discussion throughout the work. Without his advice and encouragement, this work would never be fulfilled. I also express my heartfelt respect to Associate Professor, Dr. P C Debenarh, Department of Orthopedic & Trauma Surgery, Shaheed Suhrawardy Medical College and Hospital.

I would also like to express my profoundest gratitude to my project advisory board / working team members; Associate Prof. Dr Sheikh Abbasuddin Ahmed, Associate Prof Dr. Kazi Shamim uzaman, Dr. Zia Uddin, consultant, Dr. SubirHossainShuvuro, assistant registrar, Dr. Abdul Hannan of Orthopedic & Traumatology Department, Shaheed Suhrawardy Medical College and Hospital.

My deepest admiration and sincerest love to and my laser machine sponsor- Sinha Abu Khalid, technician Mr. Mamun, laser operator Ms. Jannant & Ms. Chewty of LabNucleon, and Md. Abdul Aziz, sponsor- Sinha Abu Khalid, technician Mr. Mamun, laser operator Mr. Polash, Mr. Abul- kasem, Ms. Fatema, Ms. Farida, Mr. Malek of Orthopedic and Traumatology Department, Shaheed Suhrawardy Medical College and Hospital.

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   b) Department of Dermatology, Harvard Medical School
   c) Harvard-MIT Division of Health Sciences andTechnology
   d) Graduate Program in Cell Molecular and Developmental Biology, Sackler School ofGraduate Biomedical Sciences, Tufts University School of Medicine.


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