

Photodynamic Therapy: Role in Dentistry (A Brief Review)

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Abstract:

Bacteria form a biofilm that is the safest habitat for them. Pathogenic bacteria are the main etiological factor responsible for the failure of various treatments. Quorum sensing helps bacteria to communicate with each other and coordinates with their activity. Mechanical debridement with antimicrobial therapy is usually a conventional mode of periodontitis patients. Another effective tool is Photodynamic therapy for localized and superficial infections. This article summarizes the history, general principles, applications, indications, mechanism, and adverse effects of photodynamic therapy in various fields of dentistry.

Keywords: Biofilm, Antimicrobial therapy, Periodontitis, Photodynamic therapy.

Introduction

It has been known since the 20th century that visible light has damaging action on microorganisms in the presence of the dye. More recently, Photodynamic Therapy has been introduced for its anti-microbial action. It is a non-invasive method that involves photosensitizer agents know as (photosensitizers) in the presence of oxygen. It is used in treatment for various diseases such as in the patient with antibacterial, antifungal, antiviral drug-resistant, pocket debridement and gingivitis, chronic periodontitis, aggressive periodontitis, Oral cancer, peri-implantitis, in oropharyngeal candidiasis, disinfect of the root canal, mucosal infections, treating denture stomatitis, etc. It is also known as Photoactivated disinfection, Photoactivated chemotherapy.¹

It involves three components:

1. Photosensitizers
2. Light
3. Oxygen

Photosensitizers: First-generation Photosensitizer was available in the 1970s. They may be ingested orally, applied topically, or injected intravenously depending upon the agents. They possess photo-physical, chemical, and biological characteristics. Their ideal characteristics are similar to the pure form of chemical composition.²They have been classified based on their **chemical structures** are:

Tetrapyrroles: chlorophyll, porphyrins, phthalocyanines

Furocoumarins: xanthotoxin, bergaptene, psoralen, and its methoxyderivatives.

Tricyclic dyes: methylene blue, proflavine, acridine orange, erythrosine

They affect gram-positive bacteria by bearing a positive charge.

Some commercially available are Photofrin, Foscan, ALA.

Light sources: Diode laser system is commonly used since it is easy to handle, cost-effective and portable. For the treatment of large areas, sources such as tungsten filament, xenon, quartz halogen are used which are non-coherent by nature.³ LEDs are being used because they are small, lightweight, and less expensive than typical light sources. They are mostly activated by red, blue light with a depth of penetration from 0.5cm to at least 1.5cm.

Mostly Fiber optic catheters are used. Light source delivery depends upon its location and morphology of the affected site.

History

It was discovered accidentally at the beginning of the 20th century. It was applied in the medical field for the inactivation of cells, micro-organism by light activation. The term photodynamic action was introduced in 1904 by Von Tappeiner, one of the photobiology. In 1911, Hausmann conducted a study on the biological effects of hematoporphyrin and reported on the effect of light and hematoporphyrin in red blood cells and describe skin reactions in mice that are exposed to light. In 1913, the same study was carried out on humans. Later Dougherlg and Marcus renamed it “Photodynamic therapy”.⁴ PDT was first approved by the Food and Drug Administration in 1999 to treat pre-cancerous skin lesions of the face and scalp.

Principle

Photodynamic therapy is based on the principle that the photosensitizer or the photoactive molecule binds with the target cell and can be activated with the light of a suitable wavelength. This process generates free radicals that act on the cell membrane of the microbial cell leading to its toxic effects. However, it has been less effective against gram-negative as compared to gram-positive due to their special cell wall.⁷The production of singlet oxygen is responsible for reacting with biological systems and destroying them. The energy for the excited state is about 220 kcal/mol above the ground state level.

Mechanism of Action

Before continuing the therapy, protective glasses should be worn by the patient, the assistant, and the operator.

Photosensitizer or its metabolic precursor is administered to the patient.

Most photosensitizers are activated by light between 630–700nm.

Photosensitizer gets activated by two mechanisms:

Type-I: By **transfer** of hydrogen/electron directly from photosensitizer producing ions or by removal of electron forming free radicals. These free radicals react with oxygen forming highly reactive species.

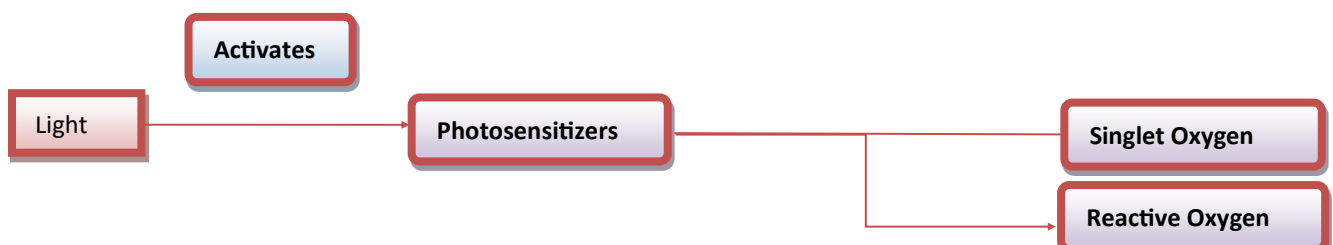
Type-II: It includes the use of triplet state photosensitizer which reacts with oxygen to form highly reactive oxygen known as singlet oxygen. This singlet oxygen interacts with bacteria to induce oxidative damage on the cell wall or cell membrane.⁵

However, the Type-II method is the most accepted pathway for bacterial destruction as it is limited to a localized area without causing any destruction to adjacent area.

Upon irradiation with light of a specific wavelength, the photosensitizer undergoes a transition from a low energy ground state to the excited singlet state.

The photosensitizer may decay back to its ground state or may undergo a transition to a higher energy triplet state.⁶

This triplet state reacts with oxygen to generate singlet oxygen and free radicals leading to toxic effects on bacteria or tumor cells.



Application in Photodynamic therapy in various fields of dentistry

Periodontology

Scaling and root planing combined with photodynamic therapy using methylene blue leads to improvement of clinical trials.

They are useful in reducing redness, bleeding on probing and the number of *P. gingivalis* bacteria.

In the treatment of Furcation area by decreasing the need for flap procedure and reducing treatment time.

It also detoxifies endotoxins such as lipopolysaccharide.

These lipopolysaccharide treated with this therapy does not stimulate the production of cytokines. Thus inactivates endotoxins. The photosensitizer compounds are topically applied in the gingival sulcus to destroy microbial pathogens.⁸

Peri-implantitis a localized inflammatory diseases affected bone and soft tissue affecting dental implant. Photodynamic Therapy helps in the treatment of decontamination to treat peri-implantitis.⁹

Oral Maxillofacial Surgery

1. Useful in the diagnosis of pre-malignant and malignant lesions.
2. The lesion healed rapidly without any acute side-effects.
3. It has been successfully used in the treatment of infections like oral candidiasis, leukoplakia.
4. Use as an effective treatment of early stage tumors.
5. It is also used in the treatment and prevention of alveolar osteitis and post-extraction pain.¹⁰
6. It also helps in the treatment of herpes labialis. It also decreases the chances of the recurrence of vesicles.
7. Photodynamic Therapy along with laser treatment helps in surgical opening of implant site for cleaning and disinfection of local defects.¹¹

Endodontics

Treatment of root canal includes three main steps access cavity, biomechanical preparation and obturation. Mechanical debridement of the canal helps to eliminate bacteria making the canal completely sterilized. However, reinfection occurs when complete elimination of bacteria is not there. This interferes with the healing of the tissue. Photodynamic Therapy along with chemo-mechanical debridement helps in eliminating chances for re-infection. It has shown promising results against endodontic bacteria.¹²Treatment of cases with chronic periapical periodontitis with photodynamic therapy is also very successful.

In Photodynamic Antimicrobial chemotherapy:

Caries results due to an imbalance between demineralization and remineralisation. Bacteria from dental plaque releases acid which is responsible for demineralization. Photodynamic therapy thus helps in the removal of dental plaque as prevention for dental caries.

Advantages¹³:

1. Resistance to treatment does not develop with repeated treatment.
2. Has moderate side effects
3. Non –invasive technique

4. No need for anesthesia
5. Can target accurately or precisely in localized sites
6. Healing is without or little scarring
7. Improve the quality of life and lengthened survival
8. No damage to adjunct tissue
9. Shorter treatment tissue
10. Less or no need for flap raise

Disadvantages¹⁴:

Despite having so many advantages, it still has some disadvantages:

1. Cannot be used in patients with blood diseases such as porphyria.
2. Less effective in treating large tumors
3. Not effective against metastatic tumors
4. Pain relief medication is mandatory before or after the treatment.

Side effects¹⁵:

Its side effects are divided into two categories related to the effects of light energy and the photochemical reaction.

1. Skin photosensitivity that can persist for weeks
2. Allergic reactions
3. Thermogenesis can occur if high-level lasers are used
4. Drugs make skin and eye sensitivity to light, the patient should avoid direct sunlight.
5. Photosensitizer can cause burns, swelling, scarring.

Future trends

Various in-vivo and in-vitro studies have been conducted and it has been found that photodynamic therapy is successful in various endodontic and periodontal problems. Peri-implantitis, mucosal infections, caries are their potential targets. However, Photodynamic therapy does not eliminate the need for chemotherapy but tends to accelerate the duration for treatment with low cost. Further studies are required to develop new photosensitizers, decrease the duration of photosensitivity, and effective light delivery systems so as decrease their side effects. Recent advances include a device based on LED technology allows for the production of light inside the target tissue. This technology expand the use of Photodynamic Therapy for the treatment of moderate and large volume of refractory tumors.¹⁶

Conclusion

Photodynamic therapy is a therapeutic approach with numerous advantages thus reducing long-term morbidity. Although the use of Photodynamic therapy in dentistry is very promising it is still in its infant stage. Superficial infection, bacterial and fungal infections seem to be an area that holds promise to incorporate this therapy as a treatment option. They are an excellent substitute for currently used chemotherapy. They can also be used as an alternative therapy for residual pocket treatment.¹⁷

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