

CASE REPORT

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# Multiprotocol treatment modality in chronic peri-implantitis utilizing decontaminating, single phase bio-ceramic and autologous fibrin matrix

Fernando Manuel Pinto Duarte, Fernando Luiz Duarte de Almeida, Leonel Alves de Oliveira

## ABSTRACT

**Introduction:** Peri-implant diseases are defined as pathological inflammatory reactions in the tissue surrounding osseointegrated implants. They are classified into two categories: mucositis—defined as peri-implant soft tissue inflammation and peri-implantitis—bone loss in the peri-implant region.

**Case Report:** A clinical case of a 61-year-old woman with an implant in the 46 anatomical position presented with peri-implantitis after a 5-year follow-up. A probe depth of 5 mm and a bone defect length of 35% was observed. The Implancre Protocol was applied, with the main objective of eliminating the biofilm present on the exposed implant surface. There are multiple approaches to treat peri-implant diseases. While non-surgical treatment is essentially sufficient for mucositis control, the treatment in most cases of peri-implantitis should be surgery. Regenerative bone reconstruction promotes bone repair in the defect area and reduces bleeding during probing. To achieve that, autologous fibrin combined with Cerasorb M was used.

**Conclusion:** The proposed approach in the presented clinical case involves the application of Implancre Protocol, whose combination of the physical decontamination technique, together with the use of chlorhexidine and orthophosphoric acid, added with the combination Piperacillin + Tazobactam together with hyaluronic acid, provided a base that allows to regenerate bone using platelet-rich fibrin with Cerasorb M and increase the implant survival time.

**Keywords:** Decontaminating, Peri-implantitis, Platelet-rich fibrin, Regenerative protocol

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## INTRODUCTION

The dental implants arising meant a change in the paradigm of oral rehabilitation, being currently accepted as the most effective method for treating both functional and aesthetic issues.

Peri-implant diseases are classified into two categories: mucositis and peri-implantitis. The mucositis is diagnosed by the clinical signs of inflammation in the soft tissue near the implant, with no bone loss surrounding it. It may be accompanied by erythema, bleeding, and sometimes suppuration. Some authors

propose, as diagnostic criteria, the presence of bleeding in the probe  $\geq 4$  mm in the absence of bone loss [1, 2]. Mucositis is reversible through a conservative approach with mechanical debridement and chemical treatment with chlorhexidine. Peri-implantitis involves peri-implant bone tissue in addition to the mucositis. It is diagnosed radiologically, due to the loss of bone supporting the implant, accompanied by bleeding and probing depth  $\geq 5$  mm [3].

## Histology and microbiology

The microbiology of peri-implantitis is more diverse than that of periodontitis, with lower levels of complex red species. Histologically, peri-implantitis is much more infiltrative near the alveolar crest and often does not have a protective layer of tissue over the bone as we normally see in periodontitis. But the most important thing for clinicians is that these diseases do not respond to treatment in a similar way [4–6].

## Risk factors

Among the most important risk factors are the lack of oral hygiene, a history of periodontitis and tobacco consumption. Other factors, such as metabolic diseases (diabetes), alcohol consumption, genetic susceptibility, absence of keratinized mucosa, type of implant surface or occlusion can also increase the risk of peri-implantitis [3, 6]. Despite this high prevalence, treatment options have been poorly studied; a 2011 Cochrane review concluded that the evidence available on the treatment for peri-implantitis is of insufficient quality and quantity and more and better research will be needed [7].

## Treatment options

Non-surgical treatment of peri-implantitis using laser or air abrasive systems has shown insufficient results. Studies that evaluated chemotherapy and mechanical debridement show minimal resolution. The attempt to use photodynamic therapy to treat peri-implantitis was also unsuccessful. We can therefore say that non-surgical treatments are not able to stop the progression of the condition [8–10].

The only treatment that shows efficacy in resolving peri-implantitis appears to be surgical. However, resection surgery is only partially effective; Leonhardt et al. [11] described the effective surgical and antimicrobial treatment in just over half of peri-implantitis lesions over a five-year period. Heitz-Mayfield and collaborators [12] demonstrated that an antimicrobial protocol with access to the surgical flap was able to stop the progression of peri-implantitis in 90% of cases in the short term (one year), but probing bleeding persisted in almost 50% of those same cases. Although a surgical approach to resection seems to improve the results, it is the combination of surgical approach and regenerative procedures where there is the highest success rate. Schwarz et al. [13] found

that regenerative surgical treatment is effective over two years, resulting in the stagnation of peri-implant bone loss and in the reduction of bleeding on probing from 80% to 34%.

Unfortunately, not all peri-implant situations are conducive to regeneration. In some of these cases, the defect will present itself as a complete loss of the surrounding bone walls, leaving regeneration as an unpredictable treatment option. Aljateeli and collaborators [14] proposed a decision tree based on the defect's morphology. If the defect had enough walls (two or more), regeneration would be attempted, but if there were zero or one wall, an apical repositioning flap would be suggested with implantoplasty (removal of exposed implant grooves and regularization of the surface with rotating instruments). The theoretical benefit of implantoplasty would be the production of a surface that is less favorable to bacterial colonization and so an easier mechanical decontamination of the surface. Charalampakis and collaborators [15] evaluated the longevity and the incidence of multiple different treatments recurrence in peri-implant pathologies; more than half of the cases evaluated relapsed and were not properly controlled. This means that peri-implantitis is not only difficult to treat, but treated cases must be carefully monitored, since recurrence is common.

In all the treatments described, one of the critical steps is always the decontamination of implant surface. The complex topographies of modern surfaces offer an excellent refuge for bacterial growth, however, there are several options for decontaminating them. Anti-infectious treatments aim to detoxify the implant surface, where the following stand out: chlorhexidine, tetracycline, metronidazole, citric acid, laser and photodynamic therapy. Mechanical debridement aims to completely remove the biofilm, using: titanium, plastic or steel curettes, saline wash, cotton gauze, air abrasion, or implantation. Some clinicians choose a combination of treatments in an attempt to ensure that surface decontamination is most successful.

Several studies have evaluated the effectiveness of different surface treatments. Schwarz and collaborators [16] compared surface decontamination using laser (Er: YAG) versus plastic curettes with cotton balls soaked in saline and found no noticeable differences in the final result. Romeo and collaborators [17–19] demonstrated that implantoplasty improved the non-regenerative surgical result of peri-implantitis, reducing the probing depths from 5.5 to 3.6 mm and the average bleeding index from 2.3 to 0.5.

Implant surgery provides decontamination of the implant surface, but there are four concerns regarding implantoplasty: (a) heat production, (b) deposition of implant material in the surgical area, (c) damage to the implant surface, and (d) weakening of the implant's internal structure. Heat production is easily controlled through effective irrigation; depositions do not appear to be associated with any adverse clinical events. Removing

the micro- and macro-rough surface of the implant is commonly considered desirable, reducing the surface roughness to minimize bacterial colonization. The ultimate concern of weakening the implant's internal structure is not often discussed because the amount of surface to be removed is minimal.

## Implacure protocol

The Implacure Protocol consists of the application of an antibiotic solution of piperacillin and tazobactam in the peri-implant pocket in two sessions, separated by 4–7 days as an initial procedure. The tip of the needle should be bent to imitate a periodontal probe, and carefully the liquid inserted into the pocket, similar to the periodontal probe. When it reaches the bottom of the bag, the solution must be injected until the it is completely filled. Make sure that the entire surface of the infected implant is covered with the liquid.

Subsequently, the implant decontamination protocol is followed: full thickness flap is performed to obtain adequate access to the treatment area, if possible with removal of the implant-supported crown or prostheses, if it is screwed on. Perform a complete curettage of the infected bone, using drill no. 1 (black ring) to clean the implant surface on the cervical part and drill no. 2 (green ring) to clean the most apical grooves of the implant surface. Place a sterile compress around the implant to protect the bone walls and surrounding soft tissue; moisten with saline to improve adherence. Apply the gel composed of 37% orthophosphoric acid and 2% chlorhexidine over the entire implant surface using the syringe and let the gel act for 2 minutes to facilitate the disintegration of the biofilm. After 2 minutes, remove the gel with a sterile cannula and wash the implant surface with saline irrigation for 10 seconds, then remove the remaining saline solution with a sterile cannula and remove the compress. The next step consists of rolling the implant with a sterile compress, impregnated with the sodium hyaluronate-piperacillin-tazobactam solution; wait 5 minutes and remove the compress. Mix the bone graft Cerasorb M with the sodium piperacillin-tazobactam hyaluronate solution in a sterile container, place the bone graft in the defect and cover the area with a collagen membrane previously soaked with the sodium piperacillin-tazobactam hyaluronate solution and suture.

## Bone regeneration protocol

The Cerasorb M is a resorbable and pure phase beta-tricalciumphosphate ceramics for filling, binding, and reconstruction of bone defects as well as for bone fusion in the entire skeletal system. Cerasorb M is made of biocompatible synthetic ceramic material with a purity of  $\geq 99\%$  and nanostructured porosity (Figure 1).

In the Cerasorb M presentation the granules have a polygonal shape and, due to the open intercommunicating multiporosity, composed of micro-, meso-, and

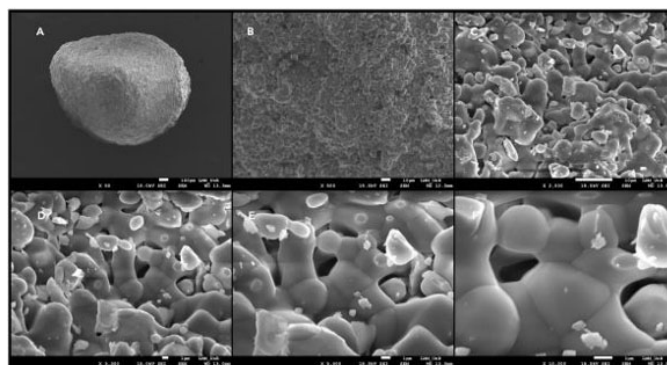


Figure 1: Cerasorb M granular units have the following structural characteristics: (A) Regular granulations with rounded edges in a spherical shape; (B–F) Architectural micro- and nanoporosities characterized as intrinsic granule porosity. These structural attributes are perceptible by ultrastructural analysis by electron micrographs, with magnifications from 50 to 10,000 $\times$  were obtained by scanning electron microscopy at the Laboratory of Microscopy and Microanalysis of the Institute of Biology at the University of Brasilia. To obtain the images, the samples were placed on a copper support, metallized and taken to the JSM 7001-F electron microscope (JEOL, Japan).

macropores (about 65%), the radiopacity is lower and absorption is faster. Such properties provide strategic anchoring points for cellular filopodia (projections of spreading membranes for cell interaction/signaling with the three-dimensional structure of the medium). Over months, in contact with the vital bone, the material is resorbed by the body and simultaneously replaced by autologous bone tissue.

As a synthetic and bioactive ceramic material, Cerasorb M has excellent histocompatibility and has no local or systemic toxicity. Unlike materials of biological origin, Cerasorb M presents no risk of infection or allergic reaction.

## Fibrin system protocol

The platelet-rich fibrin is a therapeutic blood matrix obtained by selective centrifugation acting as an adjuvant in tissue repair. For fibrin matrices obtaining, six samples of autologous blood were collected in 10 mL pure glass dry tube (Montserrat, Brazil) and two blood samples in polyethylene dry tube (Greiner Bio-One, Austria), both centrifuged in the Fibrin System centrifuge (Ortoalresa, Spain) according to the methodological proposal of Oliveira et al. 2020 [20, 21] that uses the relative centrifugal force (RCF) of 200 $\times g$  for 10 minutes, for obtaining two physical forms of fibrin, polymeric or solid gel form and monomeric or temporary liquid phase form in a single spin step.

The liquid phase was used to obtain sticky bone with Cerasorb M. The autologous fibrin membranes were applied as a recover surgical site.



## CASE REPORT

A 61-year-old female patient who attended to our consultation, reporting painful symptoms at the level of the implant in the 4.6 anatomical position. The implant had been placed about 5 years before. The probing depth was greater than 5 mm. Complementary diagnostic exam (periapical X-ray) was performed, which allowed to see a significant bone loss of about 35% of the implant length. As treatment plan it was proposed to perform a surgical approach treatment combining the protocols: Implacure, Fibrin System, and Peri-implant Bone Regeneration combined with Cerasorb M (Curasan). The patient underwent systemic antibiotic, analgesic, and anti-inflammatory therapy for 8 days. After 12 months of follow-up, the patient has a favorable clinical aspect, and a good recovery of bone trabeculation is visible radiologically. The patient has no symptoms (Figure 2).

## DISCUSSION

Peri-implant mucositis affects 80% of patients and 50% of implants, while peri-implantitis affects between 28% and 56% of patients and 12–43% of implants [19]. Both pathologies have an infectious nature as a result of the accumulation of a biofilm. The main objective of treating peri-implantitis is to eliminate the biofilm from the implant surface; in this way the progression of the infection is stagnant and the function of the implant is maintained.

Resective surgery consists of removing soft tissues and the peri-implant pocket; and may be accompanied by implantoplasty, which consists of polishing the implant surface with a diamond drill. It has been shown that this procedure favors better results than if only soft tissue resections were performed. However, this treatment is discouraged in areas of aesthetic impairment, as it does not present favorable aesthetic results. Some studies have shown that the resective treatment can recover an average of 1.9 mm of bone in the area of the defect. In addition, by promoting biofilm control in the peri-implant area, the long-term stability of soft and hard tissues is favored [18].

Regenerative surgery requires prior removal of granulation tissue from the peri-implant pocket and decontamination of the implant surface. In the literature, several antiseptics are proposed, such as 0.12 or 0.2 chlorhexidine; 3% hydrogen peroxide, sodium hypochlorite, or saline. The use of several antibiotics is also proposed: amoxicillin/metronidazole, tetracycline/doxycycline, or ciprofloxacin. The use of diode laser or Erbyum-YAG laser is also proposed to achieve implant decontamination [17]. In the present clinical case, it was decided to treat peri-implant infection by intrasulcular irrigation of piperacillin/tazobactam 100/12.5 mg seven days before the intervention, in addition to the intraoperative application of orthophosphoric acid

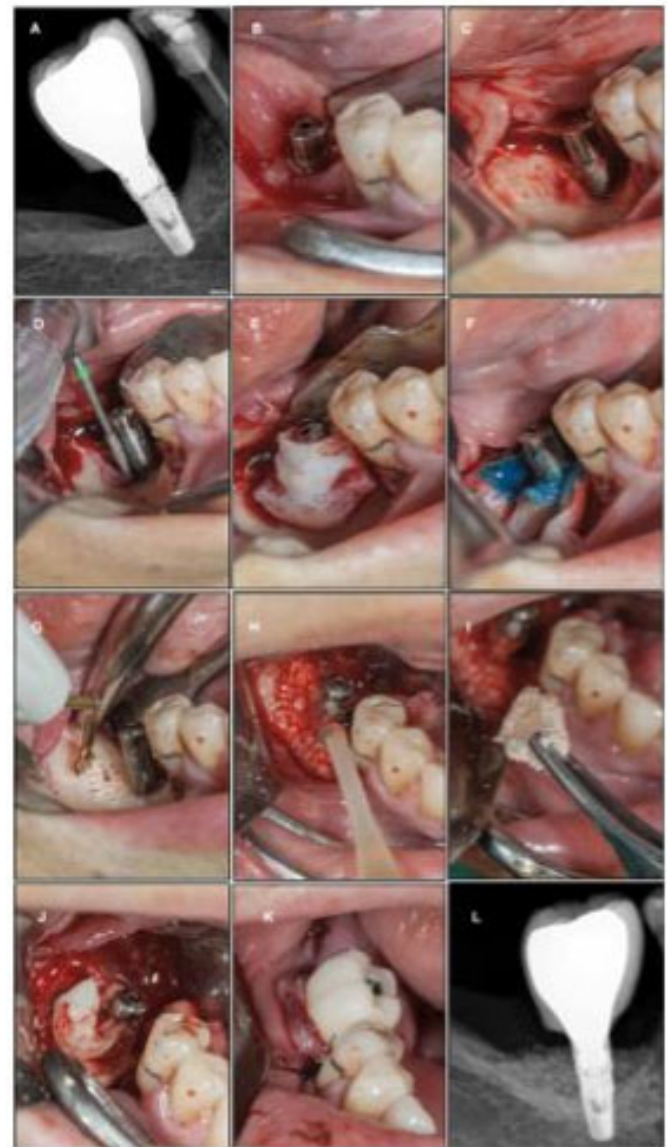


Figure 2: The sequence of image shows (A) bone loss on initial radiography; (B) Removal of the crown and installation of a healing device to protect the prosthetic connection zone of the implant; (C) Surgical flap showing bone loss and exposure implant cervical zone; (D) Exposed surface removal with diamond cutting; (E) Bone bed protection; (F) Orthophosphoric acid— $H_3PO_4$ ; (G) Piezosurgery decorticalization reaching the medullary bone; (H and I) Monomeric phase fibrin dripping to generate a means of continuity and adhesion with the sticky bone; (J) Polymeric fibrin membranes covering the graft material; (K) Temporary crown; (L) Radiograph control with 12-month follow-up.

combined with chlorhexidine and again the solution piperacillin/tazobactam [18].

Once the implant surface has been decontaminated, there are numerous materials that can be used for the regeneration of the defect: xenografts, autografts, allografts, fluoridiazapatites, and titanium granules. Studies have not shown the effectiveness of any of these regeneration materials over others [18]. In the present

clinical case, the granules of the synthetic hydroxyapatite (Cerasorb M) hydrated with a solution of piperacillin/tazobactam was mixed with liquid phase platelet-rich fibrin [20]. The objective was to promote the initial stability and the continuous release of antibiotics for the biomaterial and the environment [19].

The use of autologous fibrin in monomeric phase, in addition to serving as a binder to stabilize Cerasorb particles in the form of sticky bone, acted as a means of continuity between the medullary bone and the disinfected surface of the implant. The three-dimensional arrangement of the fibrin matrix favors cell transit allowing the progenitor cells from the bone marrow to be integrated into the graft set, constituting the tripod of tissue engineering [20, 21].

In the clinical case presented, there was a 4 mm reduction in the probing depth and no bleeding while probing, after one year follow-up.

It is essential to establish a maintenance protocol for the implants through frequent checkups and a non-surgical approach to prevent the appearance of peri-implantitis. Once peri-implantitis is established, non-surgical treatment is not effective. The type of defect must be correctly diagnosed in order to choose the appropriate surgical protocol for each clinical case.

Mechanical and chemical decontamination seems to play an extremely important role in preventing the progression of this pathology [22].

Regenerative treatment promotes bone recovery in the defect area and reduces periodontal bleeding on probing.

The approach proposed in the presented clinical case presupposes a combined treatment of implantoplasty, chemical decontamination, and regenerative treatment that showed better results when compared with those in the literature; however, it is necessary to conduct multicentric clinical studies with more significant samples and longer follow-up periods.

## CONCLUSION

It is essential to establish a maintenance protocol for the implants through frequent checkups and a non-surgical approach to prevent the appearance of peri-implantitis. Once peri-implantitis is established, non-surgical treatment is not effective. The type of defect must be correctly diagnosed in order to choose the appropriate surgical protocol for each clinical case.

## PREPRINT

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## Author Contributions

Fernando Manuel Pinto Duarte – Conception of the work, Design of the work, Acquisition of data, Drafting the work, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that

questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

Fernando Luiz Duarte de Almeida – Interpretation of data, Revising the work critically for important intellectual content, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

Leonel Alves de Oliveira – Conception of the work, Design of the work, Acquisition of data, Analysis of data, Interpretation of data, Drafting the work, Revising the work critically for important intellectual content, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

## Guarantor of Submission

The corresponding author is the guarantor of submission.

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## Consent Statement

Written informed consent was obtained from the patient for publication of this article.

## Conflict of Interest

Authors declare no conflict of interest.

## Data Availability

All relevant data are within the paper and its Supporting Information files.

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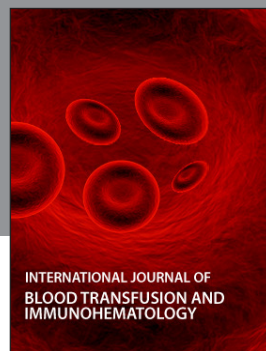
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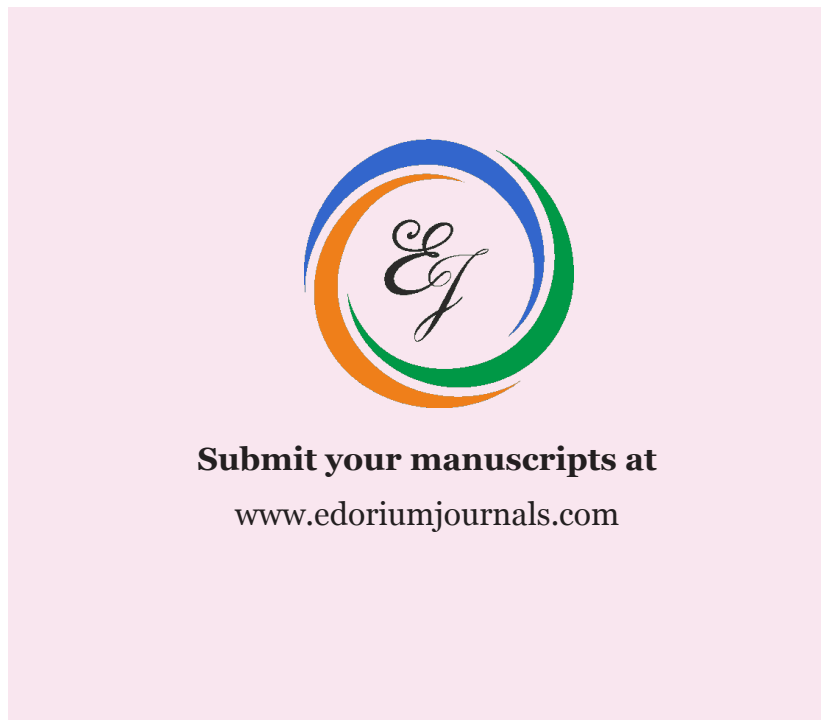
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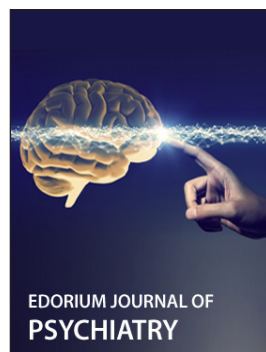
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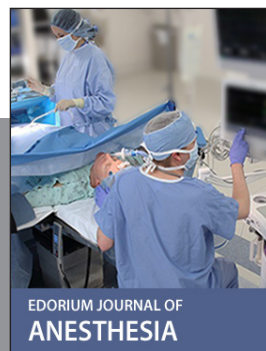
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