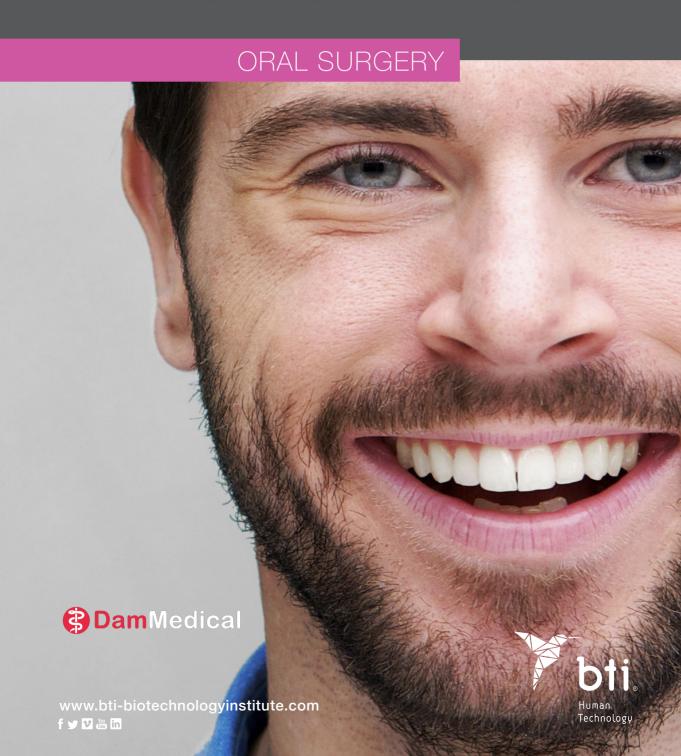


# endoret (prof.)

Endogenous Regenerative Technology



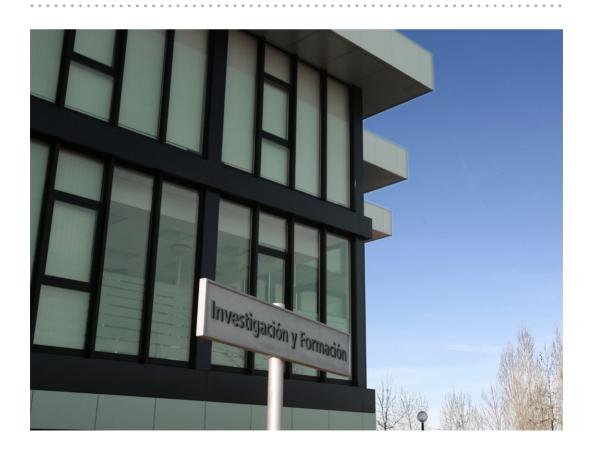


# LEADER IN REGENERATIVE MEDICINE

BTI Biotechnology Institute is a Spanish biomedicine company focused on the development of translational research projects (R&D+i).

BTI is a world-level scientific leader in regenerative medicine using Endoret in different fields of medicine.

# MORE THAN 5000 M<sup>2</sup> DEVOTED TO TRAINING, CLINICAL PRACTICE AND RESEARCH



#### WE TRAIN IN ORDER TO OPTIMISE THE CLINICAL RESULTS.

- · Specific training aimed at different medical specialities.
- More than 40 scientific collaboration agreements with universities and research institutes all over the world.
- · More than 1200 students per year of all nationalities.

# TRANSLATIONAL RESEARCH: KNOWLEDGE ACQUIRED IN THE LABORATORY APPLIED TO CLINICAL PRACTICE

• Collaboration with experts from different countries in different fields of medicine for the development of clinically effective protocols.

# MORE THAN 200 INDEXED SCIENTIFIC PUBLICATIONS BACK THE EFFECTIVENESS AND BIOSAFETY OF ENDORET®

- 20% of the workforce dedicated to research.
- · More than **15 years of research** in tissue regeneration.

## **ENDORET® TECHNOLOGY**

## 1. WHAT IS IT?

# ENDORET® IS A BIOMEDICAL TECHNOLOGY AIMED AT STIMULATING TISSUE REGENERATION BY APPLYING AUTOLOGOUS PROTEINS.

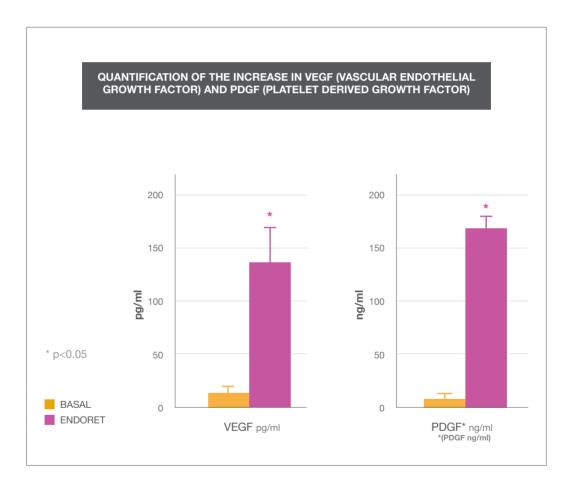
**Hundreds of endogenous proteins affect the tissue repair processes,** including angiogenesis, chemotaxis and cell proliferation. No exogenic agent can effectively govern all these processes.<sup>(1)</sup>

Endoret technology provides the necessary means for obtaining plasma rich in growth factors from whole blood.

# 2. PROTEINS AND AUTOLOGOUS MATRIX

#### A. GROWTH FACTORS

**ENDORET stimulates tissue regeneration** due to enrichment with growth factors, acting as a biological system. (2)



#### **B. FIBRIN MEMBRANE**

Enables the balanced and gradual release of a large number of molecules, including growth factors and other proteins. (3) (4) (5)

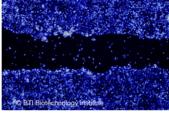
## 3. REGENERATIVE POTENTIAL

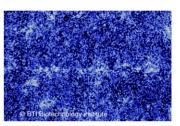
- · Promoting angiogenesis (A).<sup>(6)</sup>
- · Stimulating cell migration (B).(7)
- · Increasing the **proliferation (B).**<sup>(7)(8)(9)</sup>
- · Decreasing inflammation and pain(C).(10)(11)(12)
- · Stimulating autocrine and paracrine secretion of growth factors (B). (7)(8)(9)

(A) Pre-clinical study (B) In vitro study (C) Clinical study

#### ENDORET ACCELERATES CELL MIGRATION FOR REGENERATIVE PURPOSES (7)(8)







INITIAL STATUS (OH.)

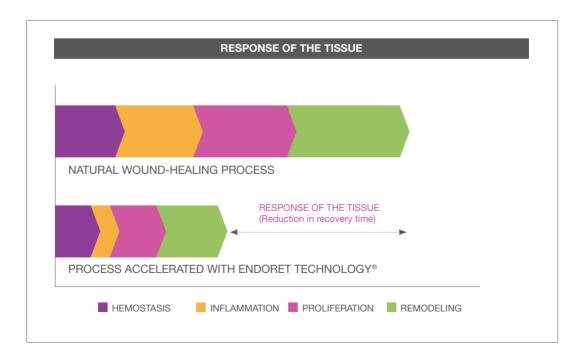
CONTROL (24H.)

ENDORET® (24H.)

# ENDORET® TECHNOLOGY REDUCES THE TISSUE REPAIR TIME IN CLINICAL TRIALS. (13) (14)

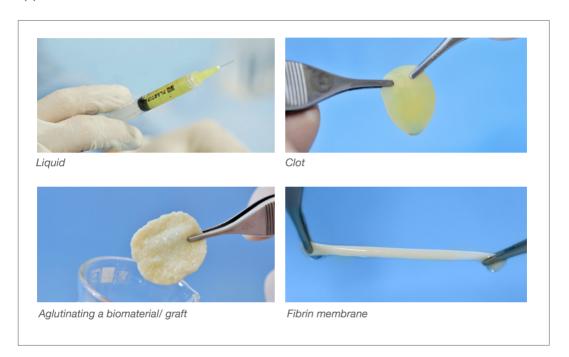


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# 4. VERSATILITY

The versatility of the Endoret technology allows for its use in different clinical applications. (15)



# 5. SAFETY

Autologous product, there are no incompatibilities nor risk of rejection.

All the formulations of ENDORET have a **bacteriostatic effect**, especially during the 4 hours after application (in vitro studies). (16) (17)(18)

More than 1,000,000 patients have been treated in more than 20 countries, without any adverse effects being reported.

### CLINICAL FEFICACY

BTI has the greatest clinical support in the world published in this field; its effectiveness is proven in more than 200 international scientific publications.

# ADVANTAGES OF ENDORET TECHNOLOGY®

Endoret is the technology with the most experience on the market in the development of specific protocols for tissue regeneration, a pioneering technology manufactured exclusively by BTI Biotechnology Institute.

### OPTIMUM CONCENTRATION OF PLATELETS

The right concentration of platelets affects the final efficacy. (2) (16)

#### FORMULATION FREE OF LEUKOCYTES

The inclusion of leukocytes increases the pain and inflammation (17) and accelerates the deterioration of the fibrin. (18)

#### CONTROLLED ACTIVATION

Enables the formation of the fibrin matrix in situ and the gradual release of growth factors, maintaining its efficacy over time. (4) (19)

#### AUTOLOGOUS

It is made from the patient's own blood, so there are no known adverse effects.

#### REPRODUCIBLE

The protocol for the preparation process and its clinical application is strictly defined and tested.

#### VERSATILE

4 preparations that can adapt the product to the clinical needs. (15)(21)

# BENEFITS AND APLICATIONS OF ENDORET TECHNOLOGY® IN IMPLANTOLOGY

### 1. INCREASED PREDICTABILITY

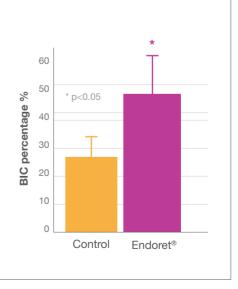
# BTI IMPLANTS COMBINED WITH ENDORET ACHIEVE A HIGH SURVIVAL RATE. (19)(20)(21)(22)

The irrigation of the implant site with ENDORET liquid, contributes to the formation of a fibrin membrane rich in growth factors on the surface of the implant, obtaining with this combination a greater apposition and bone quality. The nano-rough surface of BTI implants is specially designed to boost the biological effects of ENDORET.

HISTOMORPHOMETRIC EVALUATION OF THE BONE-IMPLANT CONTACT SURFACE (BIC) AFTER TWO MONTHS IN GOATS (23)

### SURVIVAL RATES REPORTED IN CLINICAL STUDIES:

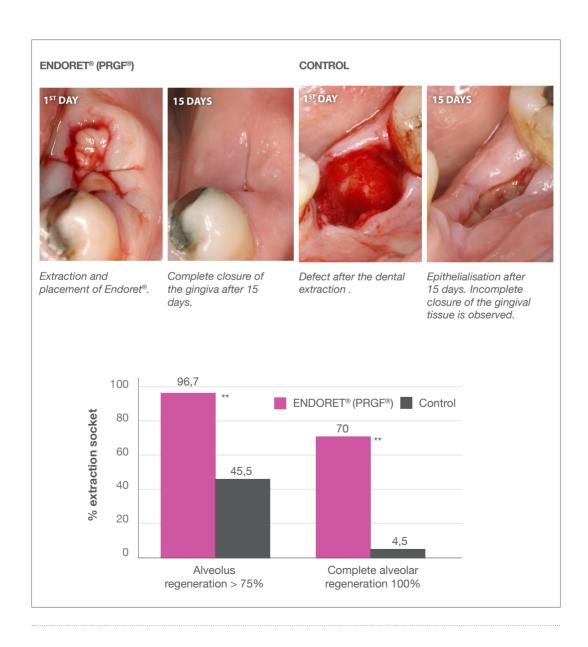
- · Up to 5 years follow-up study 5787 implants. **99.2%** (19)
- Up to 5 years follow-up study
   1139 immediate load implants. 99.3% (17)
- Up to 8 years follow-up study 1287 short implants. 99.3% (18)
- · 10-12 years follow-up study 111 short implants. **98.9%** <sup>(22)</sup>



# 2.TREATMENT OF POST-EXTRACTION SOCKET

The application of Endoret® in the treatment of post-extraction socket reduces inflammation and pain, accelerates soft tissues healing and promotes bone regeneration as it has been demostrated in several clinical trials. (11) (14) (24) (25)

The survival rate of inmediately placed dental implant, after tooth extraction, has been 98% in a clinical study. It is a safe, effective and predictable treatment. (26)



PAG. | 9 |

# 3. REDUCED RISK OF OSTEONECROSIS

The results of various clinical studies suggest that treatment with ENDORET can reduce the risk of BRONJ after a dental extraction in high-risk patients under treatment with bisphosphonates. (28)(29)

TREATMENT	NUMBER OF EXTRACTIONS	OSTEONECROSIS OF THE MAXILLA
Control	267	5
Endoret® (PRGF®)	542	0

CLINICAL TRIAL OF THE PREVENTION
OF BISPHOSPHONATE-ASSOCIATED OSTEONECROSIS
OF THE JAW (BRONJ) (28)

CONTROL

#### ENDORET® (PRGF®)









# 4. ENDORET® IN THE TREATMENT OF BRONJ

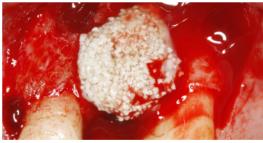
Treatment with ENDORET, after resecting the necrotic bone of BRONJ, may enhance vascularization and regeneration of bone and epithelial tissue as shown in a clinical trial. (27)

Endoret was effective in the surgical treatment of bisphosphonate-associated osteonecrosis of the jaw, achieving closure of the defect in 32 patients in a clinical prospective study. (27)

### 5. PREPARATION OF GRAFTS

ENDORET can be used to agglutinate a biomaterial, making it easier to handle and improving its osteoconductive and biological properties. (10)(30)(31)(32)(33)



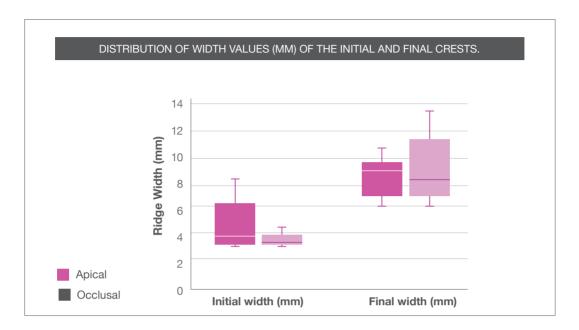


# 6. TREATMENT OF ATROPHIC MAXILLAE

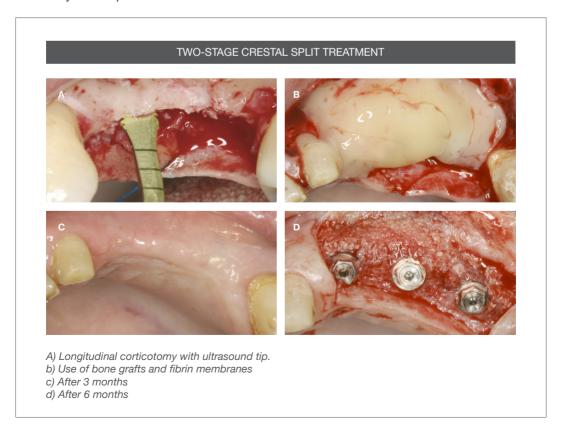
ENDORET improves the tissue regeneration and due to its versatility can be used in various surgical techniques.

#### A. LATERAL BONE AUGMENTATION

The alveolar ridge expansion and the alveolar ridge split techniques in combination with ENDORET can achieve an average bone expansion of 3.35 mm. (34)(35)



The use of ENDORET in combination with the block graft improves the tissue healing, avoiding the exposure of the graft and improves the post-operative recovery of the patient. (36)



### **B.** SINUS ELEVATION

ENDORET reduces inflammation and pain. It increases the new bone formation. (10)(32)(37) ENDORET is effective in the treatment of performations in the Schneider membrane. (31)

### SPLIT-MOUTH STUDY OF ENDORET VS. CONTROL IN A SINUS ELEVATION PROCEDURE WITH A LATERAL APPROACH. (83)



# ENDORET® INCREASES THE FORMATION OF MATURE BONE







treatment with ENDORET®

# **C.** VERTICAL BONE REGENERATION

The combination of Endoret graft with short and extra-short implants makes possible the rehabilitation of atrophic mandible without the need of more aggressive techniques. (38)(39)



# 7. PERIODONTAL REGENERATION

ENDORET may be beneficial in the field of the mucogingival surgery. (40)





Treatment of gingival defects

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