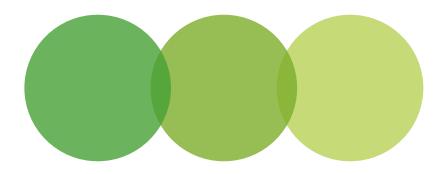
bionegen



MULTIFRACTIONAL BIO-INDUCTION

AESTHETIC MEDICAL TREATMENT FOR FACE AND BODY





REGENYAL LABORATORIES

We have been working in the field of research and manufacturing of products for the continuously evolving areas of aesthetic medicine and orthopaedics for over 20 years. This is the reason why **INNOVATION** is our mission, a mission that we pursue through the creation of cutting-edge medical devices.

We have set up an advanced manufacturing site in Spello - in the Umbria region. Our factory has certified **SAFETY** and **QUALITY** standards and is equipped with ad hoc machinery, and gives us the possibility to be even more flexible and bring onto the market innovative products controlled by highly qualified personnel along the whole manufacturing process. We are convinced that the concept of beauty cannot ignore the respect for nature, and have therefore, developed an environ-

ment friendly chain of production.



Characterized by patented and exclusive technologies, the whole production process, starting from the high quality raw material - the **ULTRA PURE HYALURONIC ACID** (provided by the laboratories of **FIDIA FARMACEUTICI**), pursues the standard of **GREEN INJECTIONS**.







MULTIFRACTIONAL BIO-INDUCTION

3 ACTIONS - 5 MOLECULAR FRACTIONS

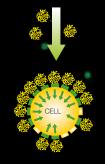
The innovative formula of Bioregen, that contains Hyaluronic Acid in fractions with different molecular weights, enables to obtain an immediate and long-lasting bio-induction of the skin, which on its turn, provides for a bio-restructuring effect from the first treatment for up to six months.



LOW MOLECULAR

WEIGHT OLIGOMERIC FRACTION

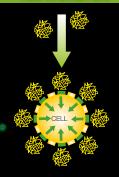
2 K DALTON



MEDIUM MOLECULAR

WEIGHT

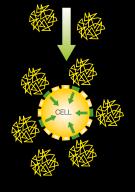
500 K DALTON
200 K DALTON
100 K DALTON



HIGH MOLECULAR

WEIGHT

1 MILLION DALTON



REGENERATION

RE-STRUCTURING

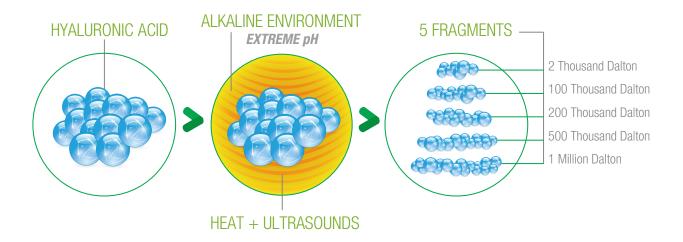
HYDRATION AND PROTECTION

STIMULATES ENDOTHE-LIAL CELLS, BOOSTS THE NEO-ANGIOGENESIS AND THE PRODUCTION OF COLLAGEN BOOSTS THE PROLIFERA-TION OF FIBROBLASTS AND BASAL CELLS OF THE EPIDERMIS

ATTRACTS AND HOLDS MOLECULES OF WATER, CAPTURES FREE RADI-CALS, BOOSTS META-BOLIC EXCHANGE

MULTIFRACTIONAL **BIO-INDUCTION**

The fragmentation



THE FRAGMENTS WITH SPECIFIC MOLECULAR WEIGHT ARE OBTAINED IN THE CONDITIONS OF **EXTREME PH**

THE FRAGMENTATION ENABLES TO OBTAIN DIFFERENT MOLECULAR FRACTIONS, EACH WITH A TARGETED ACTION. THE PROCESS IS CARRIED OUT BY IMPLEMENTING HEAT AND ULTRASOUNDS IN A SPECIAL WAY IN A HIGHLY ALKALINE ENVIRONMENT, STARTING FROM A FRACTION OF 1 M DALTON.





MULTIFRACTIONAL BIO-INDUCTION



Combats the signs of time through a natural reduction of wrinkles

PREVENTS skin ageing through hydration and stimulation, starting as early as the age of 35

PROTECTS the skin from photoageing with treatments before and after exposure to the sun

IMPROVES the quality of the areas of skin with acne-caused scars and red stretch marks

INTERVENES on the brightness of the face

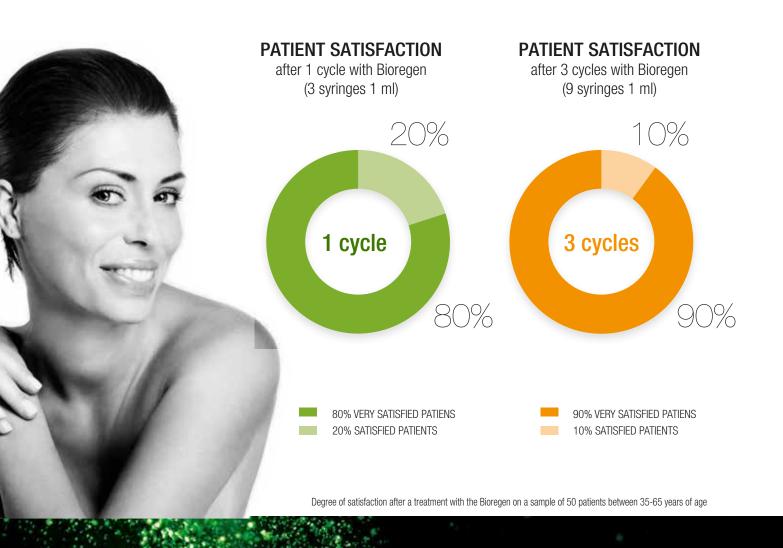
RESTORES tonus and firmness of the face and décolleté

FIRMS the skin



MAIN PROPERTIES

BIOREGEN: TECHNICAL SHEET								
COMPOSITION	DOSAGE	MOLECULAR WEIGHT	HA CONCENTRATION	PROPERTIES	BOX	CROSS-LINKING AGENT	ADMINISTRATION	RESULTS
HYALURONIC ACID SODIUM SALT IN BUFFERED PHYSIOLOGIC SOLUTION	1 ML	2 THOUSAND DALTON 100 THOUSAND DALTON 200 THOUSAND DALTON 500 THOUSAND DALTON 1 MILLION DALTON	1.6 16 MG/G OF HYALURONIC ACID SODIUM SALT IN 1ML	HYDRATION PROTECTION RE-STRUCTURING REGENERATION	3 PRE-FILLED SINGLE USE SYRINGES	NOT CROSS-LINKED	1 ML EVERY 2 WEEKS FOR 6 WEEKS AFTERWARDS 1 ML A MONTH	PREVENTS SKIN AGEING PROTECTS THE SKIN IMPROVES THE QUALITY OF THE SKIN



MULTIFRACTIONAL BIO-INDUCTION

INJECTION TECHNIQUES

Linear or micro-papular technique

BIOREGEN can be used with the linear as well as with the micro-papular technique. In the case of the periocular area it is recommended to mark single lines and to correct them always with the superficial linear technique. Small amounts of hyaluronic acid should be released while withdrawing the needle. In order to increase the aesthetic and bio-inductive effect, these injections need to be crossed with additional linear injections until reaching the zygomatic area.



SUPERFICIAL LINEAR TECHNIQUE

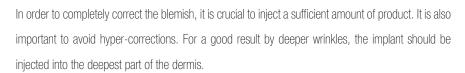
The perioral district requires superficial injections with the linear, or preferably, with a serial micropapular technique along the visible lines, and two additional linear injections into the medium dermis, parallel to the rima oris.

The needle has to inserted along the wrinkle and then gradually withdrawn releasing the product. After the injection, if necessary, the area treated should be gently massaged, in order to shape it. For the correction of fine wrinkles it is mostly recommended to inject the material into the papillary dermis.



MICRO-PAPULAR TECHNIQUE

Due to its ability to widespread homogeneously, this product is especially indicated for treatments of the neck. The linear, or the serial micro-papular, techniques provide for a good correction of the blemishes. In order to increase the bio-inductive effect it is recommended to carry out additional linear injections, parallel to the previous ones, or micro-papular spread over the area.



Bioregen can also be implemented for rejuvenation treatments on hands with micro-papular technique, followed by a gentle massage. In the case of red stretch marks the recommended technique

consists of a series of micro-injections along the stretch-mark and a slight massage after that.

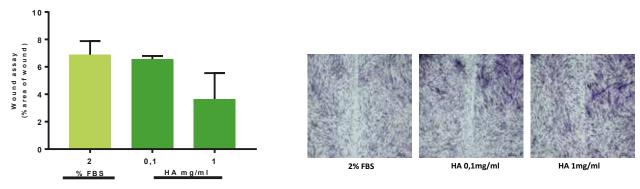


LINEAR OR MICRO-PAPULAR TECHNIQUE



MICRO-PAPULAR TECHNIQUE

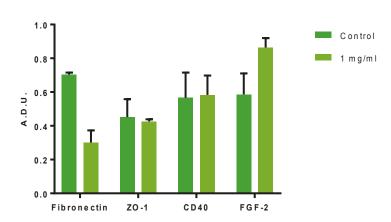
In vitro studies carried out in collaboration with the University of Siena show an efficient bio-inductive action on fibroblasts and on endothelial cells.



The cells were treated with Bioregen (in the presence of the antimitotic agent ARA-C) for 18 h. Results are expressed as percentage of wound area respect to time 0, measured by image analysis software (n=3 in triplicate). Fibroblasts try to cover the wound in the presence of Bioregen.

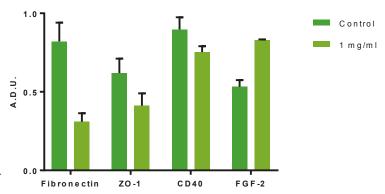
Evaluation of angiogenic and stromal markers in Fibroblasts

Cells at 90% of confluence were treated with Bioregen (1 mg/ml, 18 h). The graph is the result of n=3 experiments, and represents the arbitrary densitometric units (ADU) of the protein of interest respect to beta actin, a control protein. Fibronectin levels decrease while FGF-2 increase after Bioregen treatment of fibroblasts.



Evaluation of angiogenic and stromal markers in endothelial cells

Cells at 90% of confluence were treated with Bioregen (1 mg/ml, 18 h). The graph is the result of n=3 experiments, and represents the arbitrary densitometric units (ADU) of the protein of interest respect to beta actin, a control protein. The levels of fibronectin and ZO-1 are reduced, while FGF-2 levels are increased. The inflammatory marker CD40 does not change.



Markers Legend

Fibronectin: Extracellular matrix protein involved in cell adhesion. Its expression is reduced with cell migration.

ZO-1: Zonula occludens-1 ZO-1, also known as Tight junction protein-1, is a cell-cell junctional protein. Its expression decreases with cell migration and retraction, and hyperpermeability.

CD40: A marker of endothelial cell activation toward an inflammatory (monocyte adhesion marker) and angiogenic phenotype.

FGF-2: Fibroblast growth factor-2, angiogenic and prosurvial growth factor.





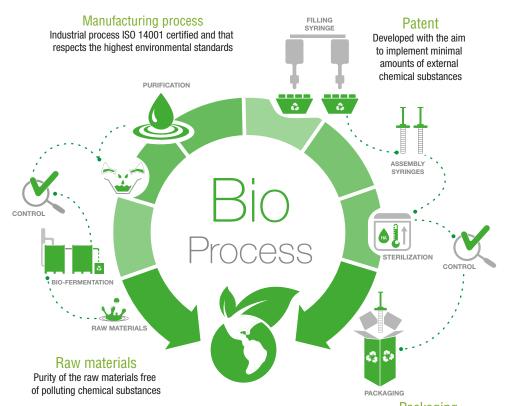
TREATMENTS IN THE MAIN AREAS OF THE FACE





Bio Process

A SPECIAL PRODUCTION METHOD CONTROLLED AND CERTIFIED



Packaging e materials used for packaging

The materials used for packaging are created from materials that respect the environment in compliance with the FSC standards

Green Injections

GREEN INJECTIONS is an international standard for injectables that applies to the safety of the product, to eco-friendliness of the manufacturing process and to the techniques aimed at obtaining natural results and maintaining natural beauty over the years. GREEN INJECTIONS are:

GREEN PRODUCT - Products of the highest quality manufactured from the purest Hyaluronic Acid with minimal presence of external chemical substances

GREEN TECHNOLOGIES - Innovative products made in Italy, in compliance with the strictest rules on the subject of environment: limited waste, energy saving, water purification, recycling, use of machinery at low energy consumption (ISO 14001 certified)

GREEN TECHNIQUES - Techniques for injections with needles or cannulae, worked out by highly qualified physicians. These techniques offer efficient and safe treatments, reducing to the minimum the risks of complications in the "difficult areas".

GREEN RESULTS - Visual "Ecologicity" of the correction that enables to obtain natural results.



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