RELIFE 5R INTEGRATED APPROACH





REshape

REboost

REstore

REsurface

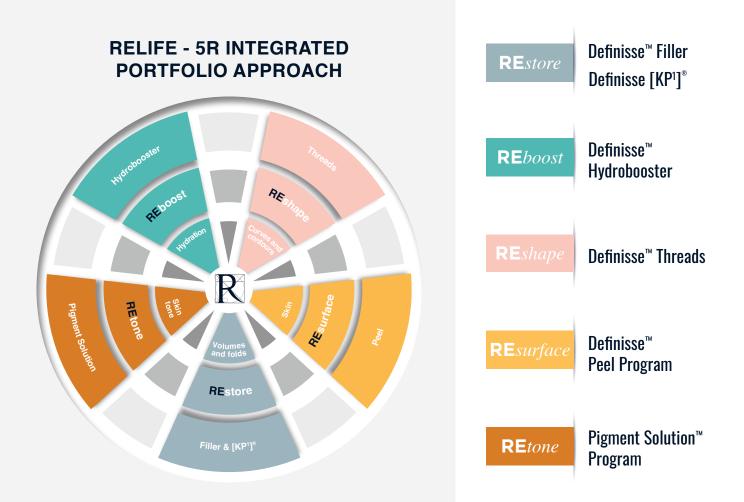
REtone

The whole is more than the sum of its parts Aristotle (384-322 BC)

Conscious Innovation in Well-Being

It is time for a new approach that considers the skin as the mirror of a complete well-being. RELIFE is a long-term project focused on helping people to achieve a complete well-being, starting from their skin. RELIFE partners with Specialists and Scientific Societies from around the world to provide consumers with high quality products based on scientific innovation.

This is why at RELIFE we talk about conscious innovation: premium, science-based innovation at the service of our consumers.





Definisse[™] Filler



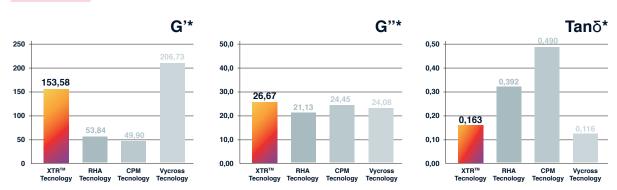


Definisse[™] touch filler

with Lidocaine

The device is intended to correct moderate-to-severe facial wrinkles and folds, and to increase lip volume.¹

Injections in the periorbital region (including tear trough, upper periorbit and eyelids), glabella and nasal alae require appropriate training.¹



Rheology²

*Rheology, 30°C at 0.7Hz. Internal Data.

HA concentration	23 mg/ml
Lidocaine concentration	0,3%
Technology	XTR [™] (eXcellent Three-dimensional Reticulation)
Composition	Cross-linked sodium hyaluronate
Effective cross-linker ratio	0,15
Depth of injection	Mid-to-deep dermis
Needle provided	2 x 27G 1⁄2"
Packaging	1 x 1 ml

Graphic elaboration from Ref. 1,3.

Clinical case



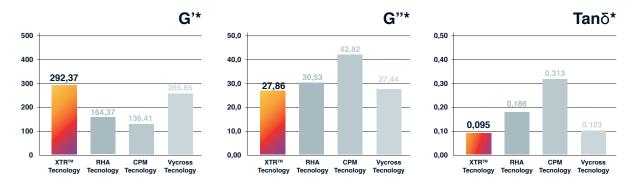
Photos courtesy of Dr. Giovanni Salti

Definisse[™] restore filler

with Lidocaine

The device is intended for correction of deeper wrinkles and folds, increase or restore facial volume and remodel facial contours.⁴

Rheology²



*Rheology, 30°C at 0.7Hz. Internal Data.

HA concentration	23 mg/ml
Lidocaine concentration	0,3%
Technology	XTR [™] (eXcellent Three-dimensional Reticulation)
Composition	Cross-linked sodium hyaluronate
Effective cross-linker ratio	0,27
Depth of injection	Deep dermis or subcutis
Needle provided	2 X 27G ½"
Packaging	1 x 1 ml

Graphic elaboration from Ref. 3,4.

Clinical case



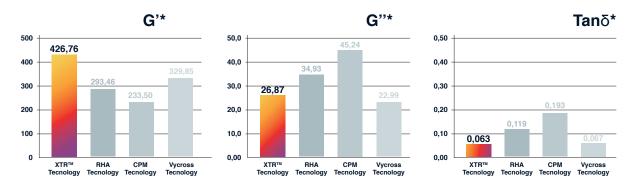
Photos courtesy of Dr. Adriano Santorelli

Definisse[™] core filler

with Lidocaine

The device is intended for **facial tissue augmentation** in order to correct midface volume deficit, for **restoring facial volume** and **shaping the contours of the face**.⁵

Rheology²



*Rheology, 30°C at 0.7Hz. Internal Data.

HA concentration	25 mg/ml
Lidocaine concentration	0,3%
Technology	XTR [™] (eXcellent Three-dimensional Reticulation)
Composition	Cross-linked sodium hyaluronate
Effective cross-linker ratio	0,49
Depth of injection	Deep dermis, subcutis, or supraperiosteal
Needle provided	2 X 27G ½"
Packaging	1 x 1 ml

Graphic elaboration from Ref. 3, 5.

Clinical case



Photos courtesy of Dr. Giovanni Salti



Definisse [KP¹][®]

Collagen Modulator Bio-peptide



DEFINISSE [KP¹][®] COLLAGEN MODULATOR WITH THE EXCLUSIVE PEPTIDE [KP¹], DERIVED FROM THE SERINE PROTEASE INHIBITOR SERPIN A1.⁶⁻⁸

Key Peptide One [KP1] showed a good collagen modulation, protecting collagen against degradation.9-12



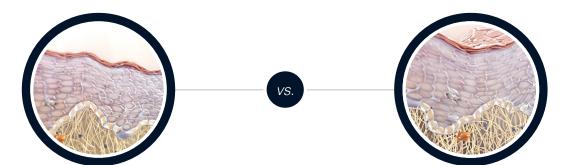
THE MAJOR MODIFICATIONS FOUND IN AGED SKIN ARE QUANTITATIVE AND STRUCTURAL CHANGES IN COLLAGEN FIBERS¹³

YOUNG SKIN

abundant, tightly packed, and wellorganized intact collagen fibrils¹³

AGED SKIN

collagen fibrils fragmented and coarsely distributed¹³

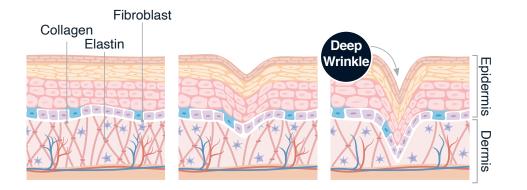


With age the skin's ability to replenish collagen 🖉 naturally

decreases by 1.0% - 1.5% per year¹⁴

THIS DECREASE IN COLLAGEN

is one of the characteristic hallmarks associated with the appearance of **FINE LINES AND DEEPER WRINKLES**¹⁴



Definisse [KP¹][®] redensifying cream

Collagen Modulator Bio-peptide



Clinically proven effective in achieving significant improvement on five anti-aging parameters after only 4 weeks of use: hydration, elasticity, density, roughness, and brightness.⁷



USER BENEFITS

The lightweight, easy-to-absorb formula provides immediate and long-lasting hydration, leaving the skin nourished, firm, and velvety smooth.⁷



ANTI-AGEING SYNERGISTIC ACTION OF:

[KP1] Collagen Modulator Bio-peptide

RELIFE's exclusive, first-in-class, bio-active peptide modulates collagen turnover by protecting it against degradation.⁷

With⁷

• Phytosphingosine, that helps to maintain the skin barrier function

- Creatine for cosmetic use, that increases cellular energy
- 2 next-generation HAs, that increase hydration and promote new collagen substrate production
 - · High molecular weight HA sits on the surface of the skin to prevent dehydration
 - Very low molecular weight HA can penetrate deeper into the skin significantly improving hydration
- Vitamin B5, stimulates the physiological skin regeneration
- Vitamin E, with antioxidant properties
- Avocado oil, that nourishes the skin

• Lactil[®], that is a blend of humectants in aqueous solution acting as a substitute of the "Natural Moisturizing Factor" (NMF) of human skin

How to use:

Apply to clean skin in the morning and evening by massaging until completely absorbed. Can be used in combination with aesthetic medicine treatments to improve results⁷

Definisse [KP¹][®] regenerating serum

Collagen Modulator Bio-peptide



Clinically proven effective in addressing dullness and reducing fine lines and wrinkles.⁸



USER BENEFITS

The highly concentrated, fast-absorbing formula provides immediate and long-lasting hydration, promoting a luminous, nourished, and velvety-smooth skin.⁸



ANTI-AGEING SYNERGISTIC ACTION OF:

[KP1] Collagen Modulator Bio-peptide

RELIFE's exclusive, first-in-class, bio-active peptide modulates collagen turnover by protecting it against degradation.⁸

With⁸

- White Truffle Extract (fragranceless), that increases skin elasticity and performs a filler effect
- 2 next-generation HAs, that increase hydration and promote new collagen substrate production
 - · High molecular weight HA sits on the surface of the skin to prevent dehydration
 - Very low molecular weight HA can penetrate deeper into the skin significantly improving hydration
- Vitamin B5, stimulates the physiological skin regeneration.

How to use:

Apply a few drops to clean skin in the morning and evening by massaging until completely absorbed. Can be used in combination with aesthetic medicine treatments to improve results.⁸

Definisse [KP¹][®] revitalising eye contour

Collagen Modulator Bio-peptide



A complete cosmetic treatment for the eye area that reduces wrinkles, dark circles, and under-eye puffiness due to an effective combination of 3 peptides and 3 latest-generation hyaluronic acids.⁹



USER BENEFITS

Lightweight and easy-to-absorb formula that provides immediate and long-lasting hydration for fresh looking skin around the eyes.⁹



ANTI-AGEING SYNERGISTIC ACTION OF:

[KP1] Collagen Modulator Bio-peptide

RELIFE's exclusive, first-in-class, bio-active peptide modulates collagen turnover by protecting it against degradation.⁹

With⁹

• **Argireline**[®], a synthetic peptide, that helps inhibit the repetitive contraction of facial expression muscles, thereby reducing hyperkinetic facial lines

- Acetil-tetrapeptide-5 proven effectiveness in reducing dark circles and under-eye puffiness
- 3 different types of HA promote new collagen substrate production:
 - High molecular weight HA sits on the surface of the skin to prevent dehydration
 - Very low molecular weight HA can penetrate deeper into the skin significantly improving hydration
 - **Cross-linked HA**, has a significantly higher ability to hydrate the skin than other HAs, and double the ability to reduce wrinkles and wrinkle depth
- Vitamin B5, stimulates the physiological skin regeneration
- Vitamin E, with antioxidant properties
- Avocado oil, that nourishes the skin

How to use:

Apply to clean skin in the morning and evening by massaging until completely absorbed. Can be used in combination with aesthetic medicine treatments to improve results.⁹

REboost

Definisse[™] Hydrobooster

The device is a sterile, absorbable, biodegradable, viscoelastic, clear, transparent, isotonic, and homogenised injectable gel implant used as a soft tissue filler without an intended medical purpose.¹⁵

The viscoelastic implant is indicated to maintain hydration, to improve tone and elasticity of the skin and, thereby, correct fine lines on the face, including lateral canthal lines (also called "crow's feet") and perioral rhytids (also called "smile lines" or "smoker's lines").¹⁵





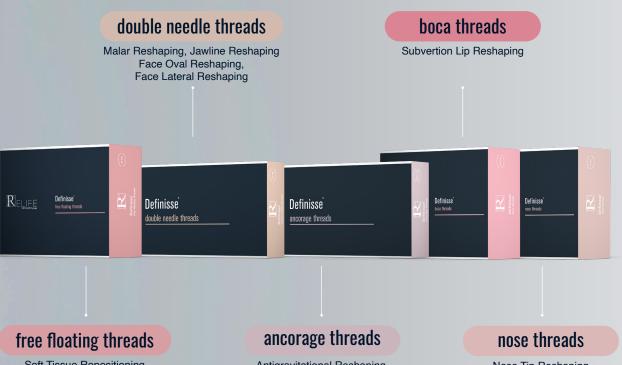
18 mg/ml + 20 mg/ml Glycerol
NA
Not cross-linked
Sodium hyaluronate + Glycerol
NA
Superficial dermal tissue
2 X 30G 1⁄2"
1 x 1 ml

Graphic elaboration from Ref. 15.

REshape

Definisse[™] Threads

Distinctive mechanical properties and effective anchoring structures. For an immediate and long-lasting facial contours reshaping.



Soft Tissue Repositioning

Antigravitational Reshaping

Nose Tip Reshaping

FOR FACIAL CURVES AND CONTOURS RESHAPING

- 1 The peculiarity of **Definisse™ Threads** is the presence of a special kind of barb, created by a unique technique with the aim of not reducing the gauge of thread¹⁶
- 2 **Definisse™ Threads** are a range of absorbable, monofilament, suspension-barbed threads of synthetic origin (poly (L-lactide-co-ε-caprolactone), p(LA-CL))¹⁶
- *3* The **Definisse™ Threads** interact with the tissues with a double action:
 - Lifting action for the support and repositioning of the tissues, thanks to the mechanical action¹⁶
 - Revitalising action stimulating synthesis of collagen, hyaluronic acid and elastin, thanks to the barbs and the specific material¹⁶

- 4 Clinical studies show that **Definisse™ Threads** allow:
 - An immediate facial reshaping¹⁷
 - A good satisfaction for the patients¹⁸
- 5 Different techniques have been developed for each reference

Definisse[™] free floating threads

The **Definisse™** free floating threads are bidirectional and convergent barbed threads.¹⁶





*The length is referred to the part with barbs



Soft tissue repositioning with 12 cm Definisse™ free floating threads



Definisse[™] double needle threads

The **Definisse™ double needle threads** are bidirectional and convergent barbed threads.¹⁶



Each box contains 6 sterile threads packaged separately

The available lengths* are¹⁶:





*The length is referred to the part with barbs

The thread ends with two straight triangular needles¹⁶:

- 100 mm long with a diameter of [0.8 0.83] mm for the 12 cm long thread
- 150 mm long with a diameter of [1.26 1.28] mm for the 23 cm long thread



Jawline reshaping with 12 cm Definisse™ double needle threads

Courtesy of Dr S. Fundarò

Definisse[™] ancorage threads

Definisse[™] ancorage threads are unidirectional barbed threads.¹⁶



Each box contains 10 sterile threads packaged separately

The available lengths* are¹⁶:

Length 12.6 cm

*The length is referred to the part with barbs



The thread ends with¹⁶:

- Straight triangular needle 152 mm, [1.26 1.28] mm diameter
- Cylindrical needle 1/2 C 31.1 mm, [0.765 0.800] mm diameter



Courtesy of Dr. Cirillo

Definisse[™] **boca threads**

The **Definisse™ boca threads** are bidirectional and divergent barbed threads.¹⁹

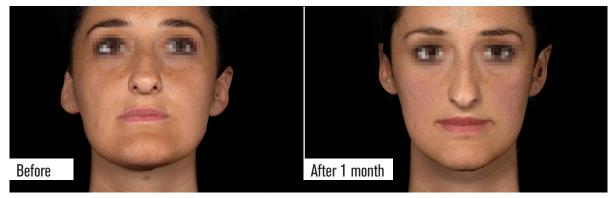


The available lengths* are¹⁹:



*The length is referred to the part with barbs





Photos courtesy of Dr. Adriano Santorelli

3D Tissue Reshaping with 9,4cm Definisse™ boca threads

Definisse[™] **nose threads**

The **Definisse™** nose threads are bidirectional and convergent barbed threads.²⁰



The available lengths* are²⁰:

Length 9 cm



*The length is referred to the part with barbs



Photos courtesy of Dr. Adriano Santorelli

3D Tissue Repositioning with 9cm Definisse™ nose threads

REsurface

Definisse[™] Peel Program

For a multilevel skin resurfacing







RELIFE RELIFE Definisse" Definisse" RELIFE tite (Persela på page site Hading pail socks i Per-Pailing Strands Definisse dry skin oily skin classic pre-peel pre-peel peel For dry, thin and For combination-oily In case of: and acne-prone skin.22 photo-ageing; superficial sensitive skin.21 actinic wrinkles or from pregnancy maintaining the skin's pH.25 expression lines; or contraceptives; post-acne skin imperfections; superficial melanotic pigmentation;

scars; seborrhoeic skin;

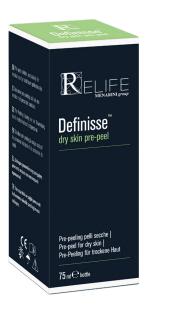
stretch marks.23

marks caused by

photo-ageing.24

RE*surface* - Definisse[™] Peel Program

Definisse[™] dry skin pre-peel





- For dry, thin and sensitive skin
- Alchool free
- It cleans gently without being aggressive (no SLES/SLES)
- It removes cosmetic product residue, excess sebum and impurities on the skin
- It evens out the skin's pH
- Key Active ingredients:
 - Lactic Acid: moisturising
 - Ethylhexyglycerin: controlling the microbial bank
 - Caprylyl Glycol: emollient²¹

MD class I

Definisse[™] oily skin pre-peel

•



Hydroalcoholic solution

- For combination-oily and acne-prone skin
- It cleans gently without being aggressive (no SLES/SLES)
- It evens out the skin's pH
- It removes cosmetic product residue, excess sebum and impurities on the skin
 - Key Active ingredients:
 - Denatured alcohol: controlling microbial bank, astringent action²²

MD class I

Definisse™ classic peel

For a multilevel skin resurfacing

Definisse[™] classic peel allows a multilevel synergic action due to the 3 main components²⁶:

2

3





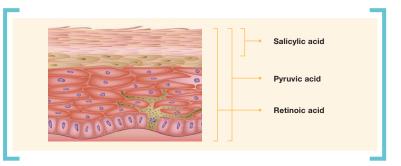
Immediate action: acts on stratum corneum via a keratolytic effect.^{23,27}

Pyruvic acid

Rapid penetration, antimicrobial and sebo-regulatory properties, collagen and elastic fiber synthesis induction, lightening action.^{26,28}

Retinoic acid

Delayed action, increased epidermic cell turn-over and collagen synthesis. $^{\rm 26,27}$



MD class I

This great synergy of action has allowed to reduce the concentration of the components, thus **reducing damage to the skin structures** treated.²³

Definisse[™] classic peel is a medical device used to promote exfoliation and renewal of the superficial layers of the skin.²³

In cases of: • superficial actinic-type wrinkles²³;

- photo-aging lesions (including wrinkles, lentigo, keratosis)²³;
- comedonal acne, post-acneic skin lesions.²³

The treatment is also a useful aid in the prevention of papular pustular acne.²³



Definisse™ lightening peel

For an additional lightening action

Definisse[™] lightening peel allows a multilevel synergic action due to the 3 main components²⁶:



address uneven pigmentation.29



Definisse[™] lightening peel is a medical device used to promote exfoliation and favour the progressive reduction of superficial dyschromia of the skin on the face and body.²⁴

For this reason, the medical device can be used to improve skin appearance.²⁴



Definisse™ neutralizing buffer





It is a neutralising buffer solution for use after a chemical peel $^{\rm 25}$

• It neutralises acids that remain on the skin after a peel, in a effective but delicate way, maintaining the skin's pH within a normal physiological range (pH 5-5.5);

• Definisse[™] neutralizing buffer does not bring the skin's pH level to basic values (which may occur when using basic neutralising solutions, such as bicarbonate of soda), therefore no extra chemical stress is exerted on the newly treated skin;

• it also contains chitosan to soothe and moisturise the skin.²⁵

REtone

Pigment Solution™ Program

A valid 1st treatment option to reduce hyperpigmentation for an even skin tone clinically proven effective and safe.

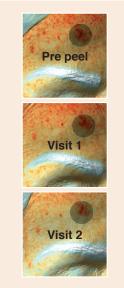
Pigment Solution Program	RELEASE		Reliefe
u dana unitarité. Mala la materi di la de norte et la que di distanta i di dina da unitari nama il la dista di la dista di la que di distanta i di dina da unitari di la di la dista di la dista di la que di di dista di la dista di la dista di la dista di la dista la dista di la parti e di la se dis sella conglegichi adhesime i à sur faro di la parti e di la se dis sellari conglegichi adhesime i à sur faro di la partir e di la se dis sellari conglegichi adhesime i à sur faro di la partir e di la se dis sellari conglegichi adhesime i à sur faro di la partir e di la se di se dista	day cream 30 mtC	properties cleaner 100mi C	night <mark>cream</mark> 30 mi C

Pigment Solution™ Program





Pigment Solution™ Program



Antera 3D[®] digital images that show the patient at different time points: pre-peel, visit 1 (post peel, enrollment, treatment) and visit 2 (end of study treatment.



Pigment[™] Solution Program

day cream

Specifically formulated to reduce hyperpigmentation.³⁰

- Cell renewal with Vitamin A³⁰
- Lightening action by inhibiting the synthesis of new melanin due to Kojic Acid³⁰
- Moisturises and restructures the skin protecting it from external agents and promotes the reduction of transepidermal water loss (TEWL) due to AquaxyI^{™30}
- Vitamin E protects the skin from various deleterious effects due to solar radiation by acting as a free-radical scavenger. Experimental studies suggest that vitamin E has photoprotective properties.³¹



Pigment[™] Solution Program preparation cleanser

 $100_{ml} e$

Gentle cleanser with a blend of soothing ingredients formulated to gently cleanse and prepare the skin for the treatment.30

Double peeling effect RELIFE **Glycolic Biodegradable Kojic** acid³² acid³² scrubbing (5.7%)(0.3%)beads (BSB)³⁰ Pigment Solution Chemical³¹ Mechanical³⁰ ght cream IN JUST Nouble-peeling depigmentation reatment | Trattamento pigmentation dark spot size epigmentante a doppio effetto **28 DAYS**: eeling | Behandlung zur epigmentierung mit doopeltern reduction:33* reduction:33* eeling-Effekt | Traitement mentant à double effet pee -14.6% -6.8%

*Test performed on 28 patients with melanin dark spots on face, hands, neckline, and chest which applied the product once a day for 28 days. Two visits were performed during the study: one at the beginning and one at the end of the treatment after 28 days.

Pigment[™] Solution Program night cream

Specifically formulated to even out pigmentation on the skin's surface 30

RELIFE is a Menarini Group Company

The Menarini Group, with its headquarters in Florence, is present in 136 countries around the world with 17,000 employees. With a turnover of 3.6 billion Euro, Menarini today is 13th in Europe out of 5.345 companies and 35th worldwide out of 21,587 (source: IQVIA). Menarini has 6 Research & Development centres and its products are present in the most important therapeutic areas including cardiology, gastroenterology, pneumology, infectious diseases, diabetology, anti-inflammatory agents, and analgesics. Menarini's products are manufactured in the Group's 16 production sites, in Italy and abroad, where more than 583 million packs are produced and distributed over five continents each year. Through its pharmaceutical products, Menarini contributes continuously and with the highest quality standards towards the health of patients all around the world.

References

- 1. Definisse[™] touch filler + Lidocaine, Information For Use.
- Salti, G., & Fundarò, S. P. (2020). Evaluation of the Rheologic and Physicochemical Properties of a Novel Hyaluronic Acid Filler Range with eXcellent Three-Dimensional Reticulation (XTR™) Technology. Polymers, 12(8), 1644.
- 3. Rigano Laboratories srl. Analysis of different types of injectable cross-linked hyaluronic acid gels. Prot. 221/19
- 4. Definisse[™] restore filler + Lidocaine, Information For Use.
- 5. Definisse[™] core filler + Lidocaine, Information For Use
- 6. Definisse [KP1]® redensifying cream leaflet.
- 7. Definisse [KP1]® regenerating serum leaflet.
- 8. Definisse [KP1]® revitalising eye contour leaflet.
- 9. Cipriani, C. et al. (2018). Serpin A1 and the modulation of type I collagen turnover: Effect of the C-terminal peptide 409-418 (SA1-III) in human dermal fibroblasts. Cell biology international, 42(10), 1340–1348.
- 10.Errante, F et al. (2020). Cosmeceutical Peptides in the Framework of Sustainable Wellness Economy. Frontiers in chemistry, 8, 572923.
- 11.Pascarella, S. et al. (2016). Serpin A1 C-Terminal Peptides as Collagen Turnover Modulators. ChemMedChem, 11(16), 1850–1855.
- 12. Rovero, P. et al. (2022). The Clinical Evidence-Based Paradigm of Topical Anti-Aging Skincare Formulations Enriched with Bio-Active Peptide SA1-III (KP1) as Collagen Modulator: From Bench to Bedside. Clinical, cosmetic and investigational dermatology, 15, 2693–2703.
- 13.Shin, J. W. et al. (2019). Molecular Mechanisms of Dermal Aging and Antiaging Approaches. International journal of molecular sciences, 20(9), 2126
- 14.Reilly, D. M., & Lozano, J. (2021). Skin collagen through the lifestages: Importance for skin health and beauty. Plast. Aesthetic Res, 8(2).
- 15.Definisse[™] hydrobooster, Information For Use.
- 16.Berardesca E. Reshaping with barbed threads. Minerva Medica. Feb 2020.

- 17.Savoia, A., Accardo, C., Vannini, F., Di Pasquale, B., & Baldi, A. (2014). Outcomes in thread lift for facial rejuvenation: a study performed with happy lift[™] revitalizing. Dermatology and therapy, 4, 103-114.
- 18.Kalyan R. et al. 2017. The Efficacy of Polycaprolactone Threads in Zygomatic and Mandibular Lifting: consecutive Study from a Single Practitioner's Experience. Int J Clin Exp Dermatol; 1(1): 1-3
- 19.Definisse[™] boca threads outer box.
- 20.Definisse[™] nose threads outer box.
- 21.Definisse[™] dry skin pre-peel Information For Use.
- 22.Definisse[™] oily skin pre-peel Information For Use.
- 23.Definisse[™] classic peel Information For Use.
- 24.Definisse[™] lightening peel Information For Use.
- 25.Definisse™ neutralizing buffer Information For Use
- 26.Cavallini M. Le iperpigmentazioni cutanee: proposta di trattamento con peeling chimico combinato e schiarente topico locale. Medicina estetica
- 27.Celleno L. Dermatologia cosmetologica. Capitolo 31- Il peeling. Pagine 362-367. Editore Tecniche Nuove
- 28. Truchuelo, M., Cerdá, P., & Fernández, L. F. (2017). Chemical Peeling: A Useful Tool in the Office. Peeling químico, una herramienta útil en la consulta. Actas dermo-sifiliograficas, 108(4), 315–322
- 29.Wong, V. (2021). Resurface, retone and reboost: a multi-modality approach for prejuvenation of millennial patients. J Clin Exp Dermatol Res. S, 8, 549.
- 30.Pigment Solution Program Information For Use
- 31.Keen, M. A., & Hassan, I. (2016). Vitamin E in dermatology. Indian dermatology online journal, 7(4), 311.
- 32.Pigment night cream labelling.
- 33.Berardesca E. et al. 2017 "Valutazione clinica e strumentale sull'efficacia di una nuova formulazione depigmentante" Dermakos, 51-53





Discover more at www.relife-icme.com



RELIFE Srl Via Dei Sette Santi 3 - 50131 Florence, ITALY Tel. +39 055 56.80.93.45 - Fax +39 055 56.80.93.46

Definisse™ Filler: CE0123 (CROMA-PHARMA GmbH), CE0459 (CROMA GmbH) Definisse™ Threads: CE0373 (ASSUT EUROPE SPA) Definisse™ Peel Program: CE Mastelli Srl

The information in this material is intended for Healthcare Professional only