







Marine bio-research, more than water

- Ocean, lifeblood of Earth, has an untapped potential, with only a 5% explored
- Unique species from the ocean's depths and sea breeze, to the marine cnidarians and onshore halophyte plants
- In collaboration with research institutes, we have proprietary collections of microorganisms for an unlimited source of new active ingredients

Over 22,000 microorganisms at our disposal





Expedition







Marine cnidarian

Halophyte Plants*

Extreme aquatic environments

...and more True novel active ingredients yet to be unveiled





Malaspina expedition, inspired by history

Inspired in the first expedition in 1789, a oceanographic research vessel set sail in a journey around the world

- ≥ 42,000 nautical miles of international waters navigated
- Over 350 marine water samples collected
- Over 120 unknown bacterial strains isolated
- Different depths, temperatures, salinities and oxygen levels









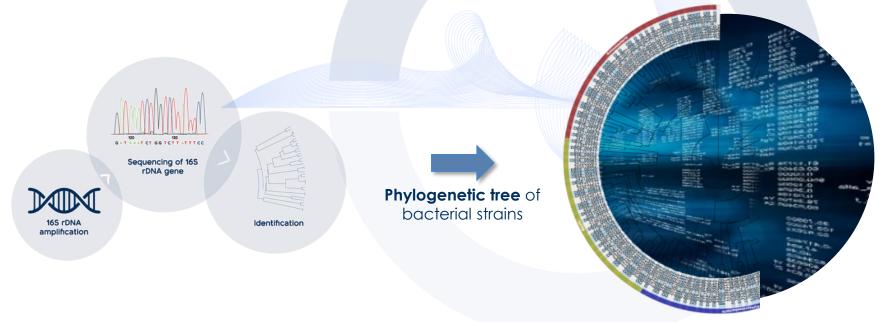
Courtesy of CSIC: Malaspina expedition, 2010





Strain classification and identification

- Classification and identification of isolated microorganisms by MALDI-TOF MS in a HTS analysis and 16S ribosomal RNA sequencing
- Cultured (only 1% of the population) and new uncultured species
- Evaluation and characterization of derived extracts through metabolomics









Transforming it into Smart Data

- Derived extracts efficacies are analyzed through transcriptomics generating a great amount of data
- Smart data for the development of new active ingredients with outstanding efficacies
- Two strategies:
 - Identify the mechanism of action of an active ingredient

Define the best active ingredient for a specific mechanism of action







Seadermium™, where fire meets the ocean



Seadermium™, is an active ingredient from a microorganism discovered during the **Malaspina** expedition, near **Reunion Island** (east coast of Madagascar)

- Young active volcanic island where burning rivers flow down the hillsides redefining the island's coastlines
- Nearby ocean currents flow, nourishing the 20 km length coral reef, filling and sustaining the rich biodiversity









The microorganism was collected at:

Depth: 3 m

Temperature: 25 °C

Oxygen level: 3.5 µmol/L



Seadermium™, nourishing premium filler

Refine your lines with SeadermiumTM to help you to bring out your true beauty, restoring skin integrity and plumping deep wrinkles

- Plumps deep wrinkles (-10% volume in 14 days, in vivo)
- Restores and strengthens skin integrity (in vitro, on human skin explants and in vivo)
- Moisturizes and smoothes mature skin (in vitro, on human skin explants and in vivo -12% wrinkles in 28 days)





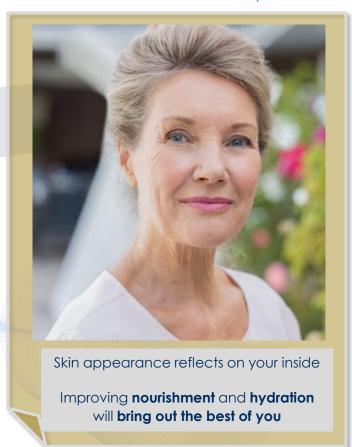


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Your skin says all of you

Book that explains what & how you have lived

- As we age, nourishment and hydration of cutaneous cells is impaired
 - Thinning of the skin and appearance of wrinkles
 - Reduction of hydration and fullness
- Skin needs nutrients to maintain its best complexion and appearance









OLD SKIN

DEJ structural integrity and cohesion

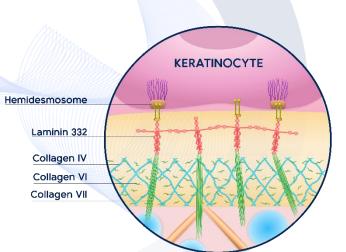
Dermis and epidermis are connected by the **dermo-epidermal junction (DEJ)**, composed of the *lamina lucida* and *lamina densa* layers

- Collagen IV, forms the 3D network of the lamina densa
- Collagen IV interacts with laminin 332 (laminin 5) and its hemidesmosomal integrin receptor



Anchoring the *lamina densa* to the deepest layer of the epidermis

- Laminin 332 has a key role in dermo-epidermal anchoring and cohesion
- Collagen VI is involved in dermal ECM → Collagen I and III



YOUNG SKIN

✓ In aged skin, there is a degradation of DEJ components, leading to a structural integrity impairment and resulting in wrinkles and sagginess





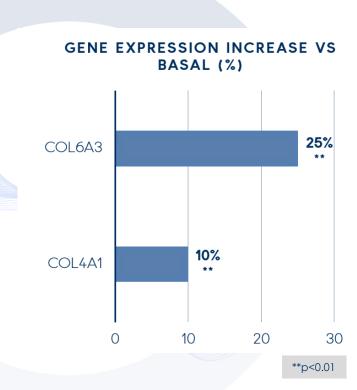
<u>In vitro efficacy</u>: transcriptomics by Dermoarray

(fluorescence)

- Primary human epidermal keratinocytes
- 0.05 mg/mL Seadermium™ active ingredient*
- 24 h incubation
- 600 genes expressed in skin cells

✓ Collagen IV and VI genes
expression increased by 10% and
25%, respectively

Up-regulation of genes involved in the dermo-epidermal structure





Microarrays can be used to study the extent to which certain genes are turned on or off in cells and tissues



In vitro efficacy: strengthening skin integrity

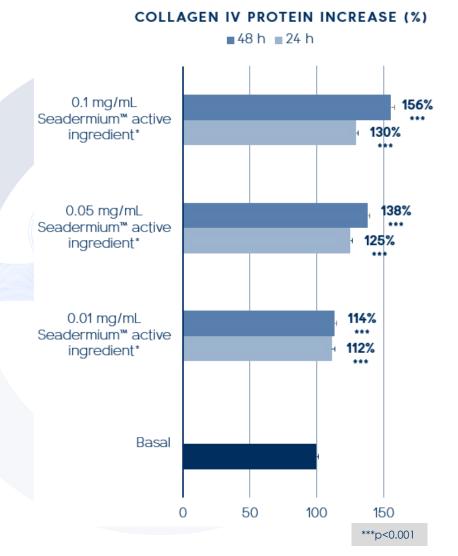


(ELISA)

- Primary human epidermal keratinocytes
- 24 and 48 h incubation
- Absorbance (450 nm)

√ 30% and 56% collagen IV increase after 24 h and 48 h with 0.1 mg/mL vs basal, recovering DEJ structural integrity

 Higher collagen IV boosting efficacy with time and dose







<u>In vitro efficacy</u>: strengthening skin integrity

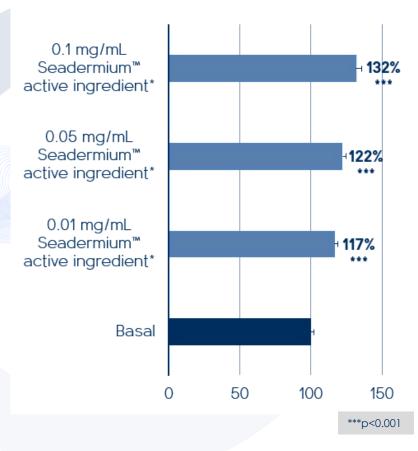


(ELISA)

- Primary human epidermal keratinocytes
- 48 h incubation
- Absorbance (450 nm)

- ✓ 32% laminin 332 increase with 0.1 mg/mL SeadermiumTM vs basal, helping to strengthen skin cohesion
 - ✓ Dose-dependent efficacy

LAMININ 332 LEVELS INCREASE (%)



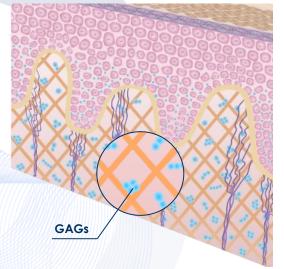






Pumping dermal water

- GAGs protect skin against excessive tissue compression and maintains its suppleness
- GAG major dermal ECM component responsible for:
 - Water-holding capacity
 - Nutrient exchange
 - Repair and protection processes
- GAGs hydrating power → helping to skin fullness



➢ GAGs levels decline with age. GAGs loss is key in the decrease of skin hydration leading to wrinkling and skin aging hallmarks





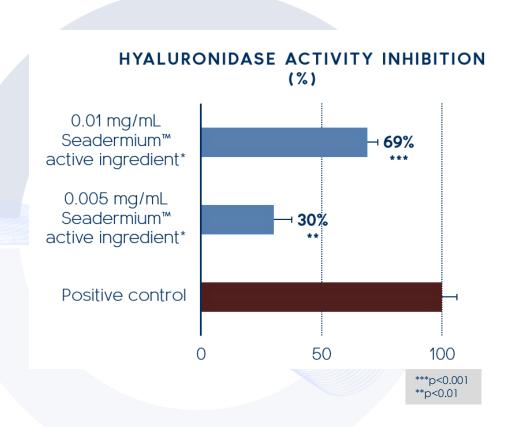


In vitro efficacy: hyaluronan protection

(enzymatic)

- Positive control
- Absorbance (570 nm)

- √ 69% hyaluronidase activity inhibition with 0.01mg/mL Seadermium™
- ✓ Protection from hyaluronan degradation helping to prevent volume loss



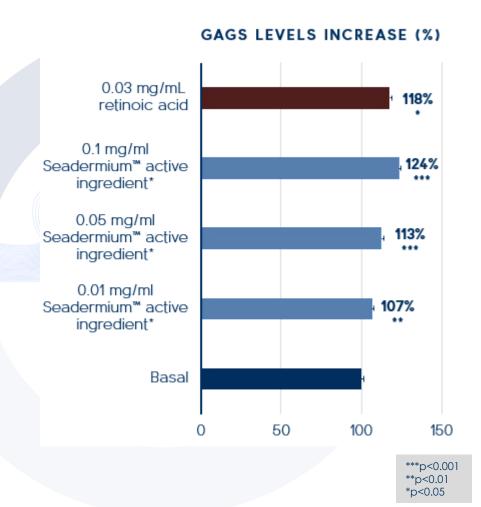




<u>In vitro efficacy</u>: hydrating volume enhancement

- Human dermal fibroblasts
- 48 h incubation
- Positive control: 0.03 mg/mL retinoic acid
- Absorbance (450 nm)

- 24% GAGs levels increase with 0.1 mg/mL, resulting in a plumping effect
- ✓ GAGs synthesis stimulation is dosedependent





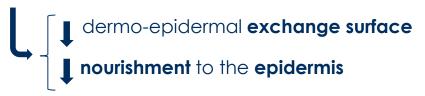


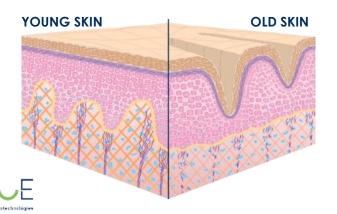
Dermo-epidermal junction, key of nourishment

 Downward epidermal ridges interdigitate with upward projections of the dermis, dermal papillae

Capillaries in the papillae provide nourishment to the epidermis

- In aged skin the number and size of the dermal papillae decrease
 - ➤ Thinner and flattened DEJ → 1 DEJ separation → wrinkle formation



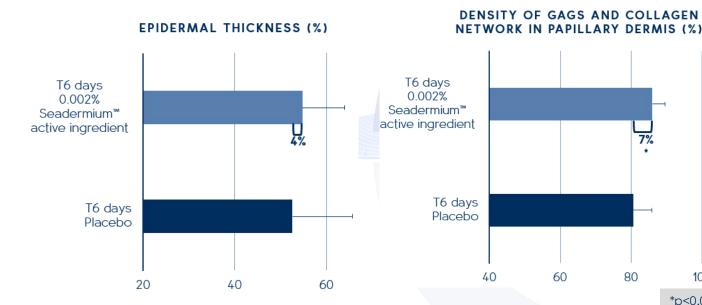


50% reduction in interdigitations per unit area of the skin between 30 and 90 years old



Human skin explants efficacy: skin replenishment and thickening (Masson's trichrome staining)

- Human skin explants (50-year old woman donor)
- 6 days
- 0.002% Seadermium™ active ingredient



√ 4% thicker epidermis vs placebo

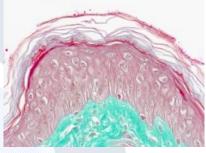
√ 7% more GAGs and collagen network density vs placebo in papillary dermis

7%

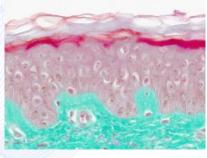
80

100

*p<0.05



Placebo



0.002% Seadermium™ active ingredient

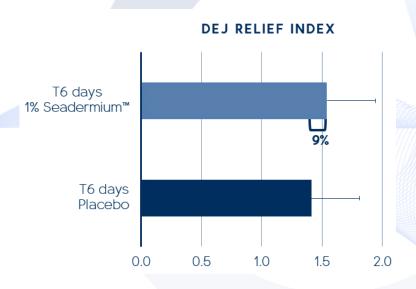




<u>Human skin explants efficacy</u>: cohesion reinforcement (immunofluorescence)

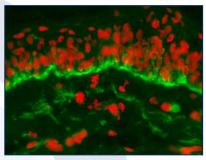
- Human skin explants (50-year old woman donor)
- 6 days
- 0.002% Seadermium[™] active ingredient

- ✓ 9% higher DEJ relief index vs placebo in 6 days (DEJ relief less flat)
- ✓ Noticeable laminin 332 increase

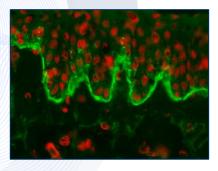








Placebo



0.002% Seadermium™ active ingredient

- DEJ relief index equals to the ratio of the length of the DEJ by the length of the epidermis.
- The flatter the relief, the closer to 1 is the DEJ relief index.

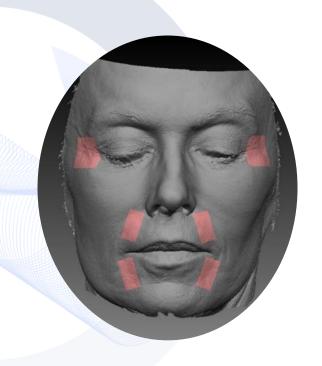




20 Caucasian women (50-70 years old) applied a cream with **4% Seadermium™** on half face, twice daily for 28 days. A placebo was applied on the other half of the face

Evaluation of:

- Type of wrinkles: Crow's feet, nasolabial fold, marionette lines (fringe projection)
- Wrinkles & smoothness (Visia®)
- Hydration (Corneometer®)







- Plumping and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 50-70





√ Excellent wrinkle plumping and smoothing efficacies

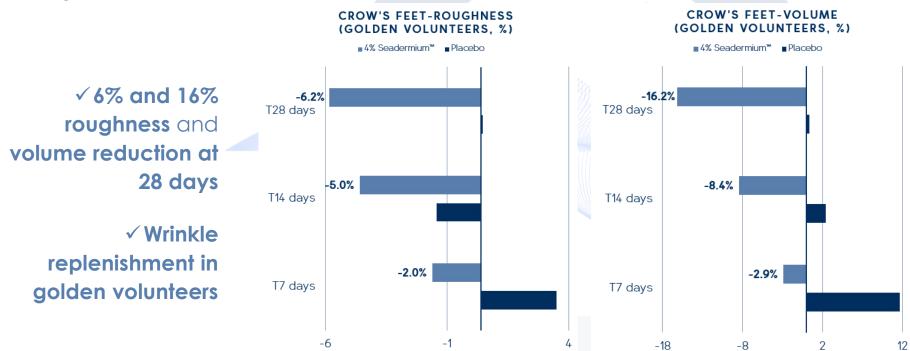






<u>In vivo efficacy</u>: wrinkle plumping & smoothness efficacy-Golden volunteers

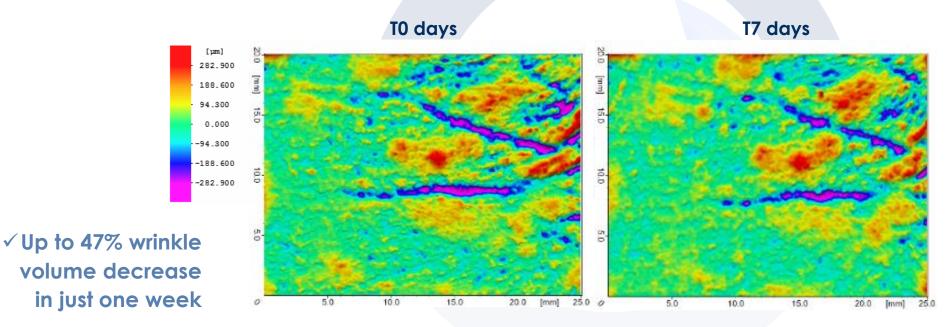
- Plumping and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 60-70







- Wrinkles and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 65 (volunteer 3)





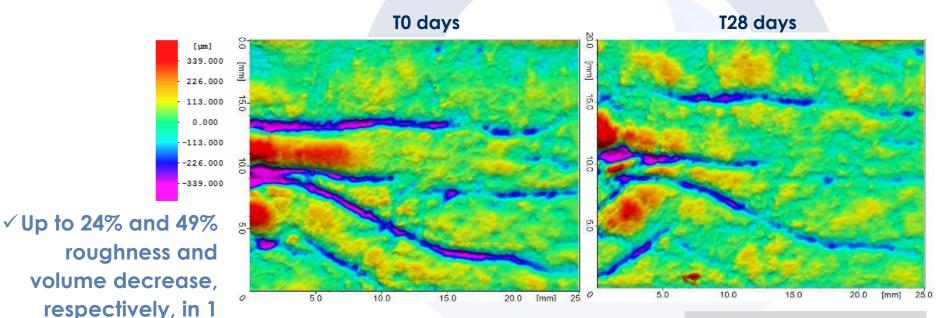
Crow's feet roughness: ↓17% Crow's feet volume: ↓47%



- Wrinkles and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily

month

Age: 67 (volunteer 20)





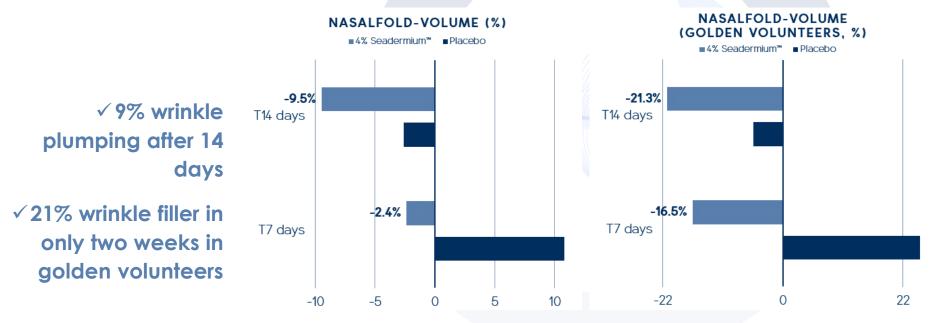
Crow's feet roughness: ↓24%

Crow's feet depth: ↓20% Crow's feet volume: ↓49%



- Plumping and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 50-70



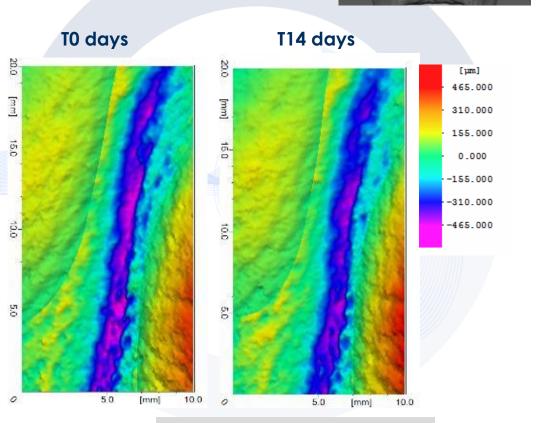






- Plumping and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 61 (volunteer 14)

✓ Up to 74% wrinkle volume decrease in 14 days





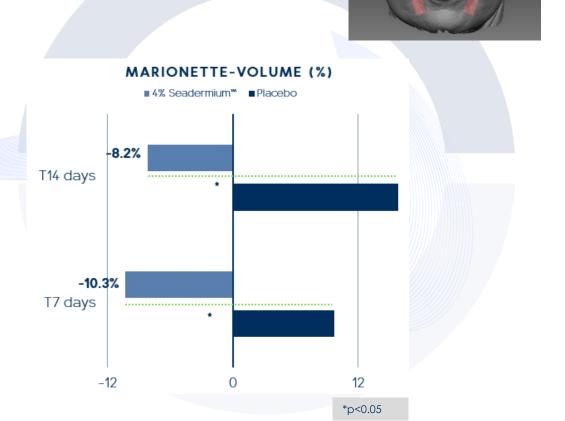
Nasolabial fold Volume: ↓74%



- Plumping and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 50-70

✓ Average 10% replenishment after only 7 days

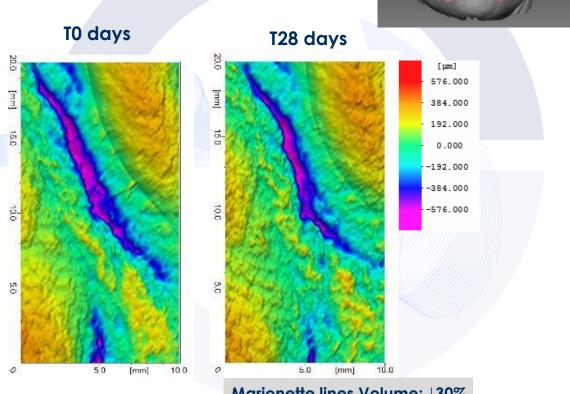
✓ Up to 58% wrinkle volume decrease in marionette lines in 28 days







- Plumping and smoothness (fringe projection)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 61 (volunteer 5)





Marionette lines Volume: ↓30%



- Plumping and smoothness (Visia®)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 50-70

√ 12% wrinkles decrease in 28 days

√ 6% smoothness improvement in 4 weeks







- Plumping and smoothness (Visia®)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 51 (volunteer 9)

T0 days

T28 days

✓ Up to 45% wrinkle reduction in 4 weeks





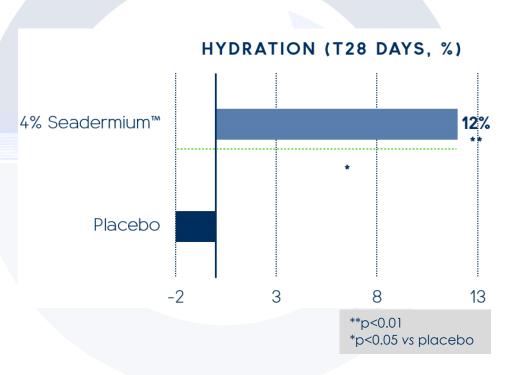
Undereye wrinkles: ↓45%





- Hydration (Corneometer®)
- Cream with 4% Seadermium™ vs placebo
- Half face, 28 days, twice daily
- Age: 50-70

✓ 12% moisture increase in one month



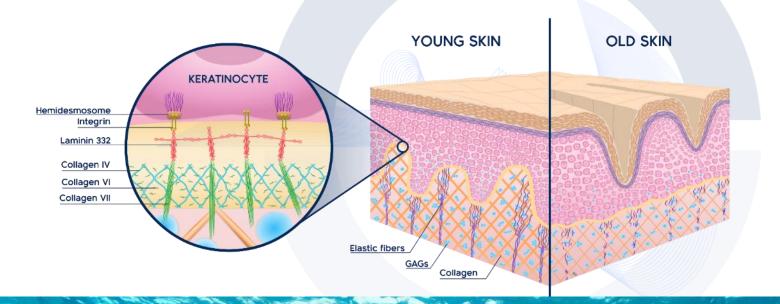






Nourishing premium filler

- Plumps deep wrinkles
- Restores and strengthens skin integrity
- Moisturizes and smoothes mature skin







Technical information

Product Name: Seadermium™

Marine active biotechnological ingredient

Recommended dose: 4%

INCI name: Bacillus Ferment



Appearance: Solution

Preservatives: None

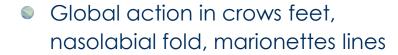
Solubility: Water soluble





Applications

Cosmetic formulations



- Targeted efficacy in golden volunteers, when you need it most
- Premium treatment for premium people
- Fill your skin with youth
- Complete approach for mature beauty















