

HYAL^o4

Skin

**Favors wound
re-epithelialization¹**



CE 0123
Medical Device

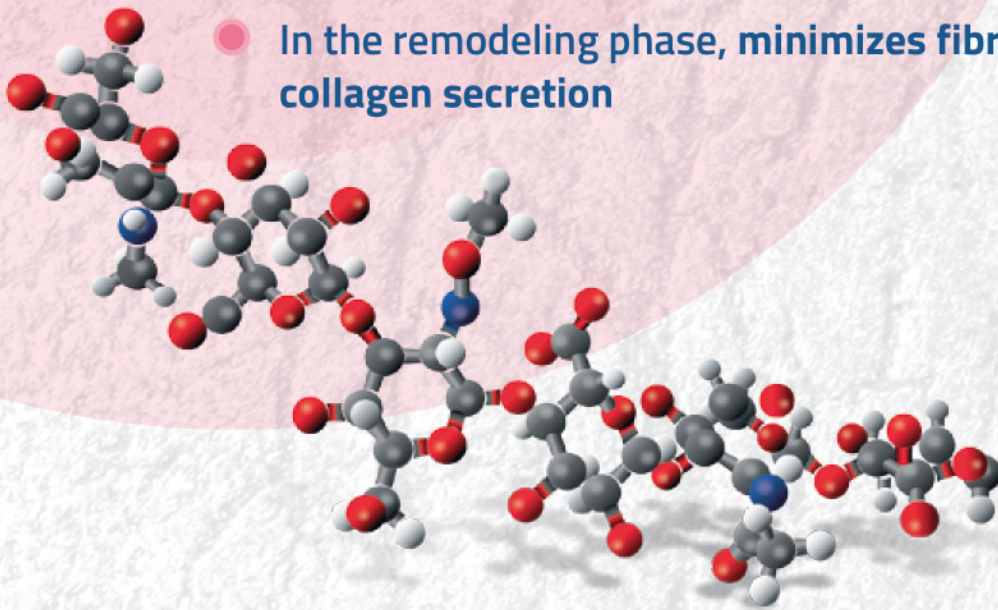
Hyaluronic Acid

**PROMOTES CELLULAR
PROCESSES ESSENTIAL
IN TISSUE REPAIR²**

Favors cell migration to ensure healing¹

LMW-Hyaluronic acid¹

- Favors tissue hydration and maintains a moist environment
- Promotes migration and proliferation of fibroblasts and endothelial cells
- In the inflammatory phase, acts as a scavenger of free radicals and activates negative feedback
- In the remodeling phase, minimizes fibrosis and stimulates collagen secretion

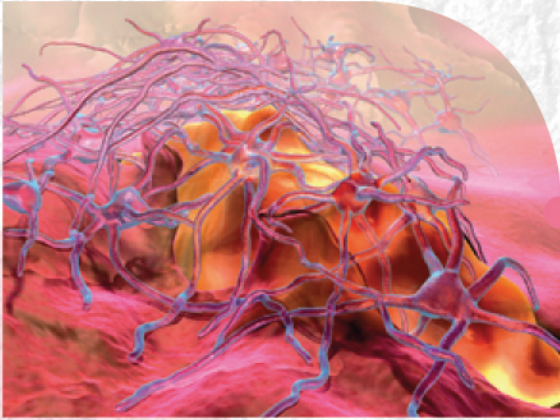


Hyaluronic acid plays a multifaceted role at each stage of wound healing³

- Increases formation of granulation tissue
- Stimulates organized deposition of collagen by fibroblasts
- Promotes neoangiogenesis
- Favors re-epithelialization

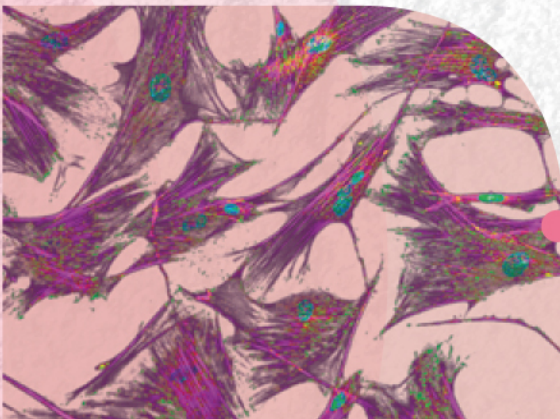
Role of HA in

Promotion of neoangiogenesis



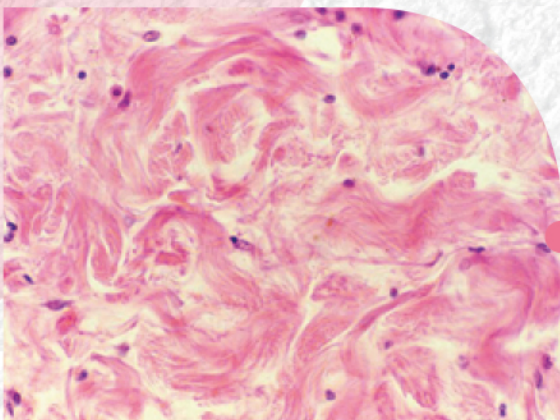
HA promotes neoangiogenesis via endothelial cell surface receptors CD44 and RHAMM. The formation of new vessels is a major step in tissue repair and wound healing.³

Migration and proliferation of fibroblasts



HA increases migration and proliferation of fibroblasts and formation of granulation tissue. Fibroblast proliferation is essential in tissue repair and HA-induced expression of TGF beta 3 may favor scarless healing.^{2,3}

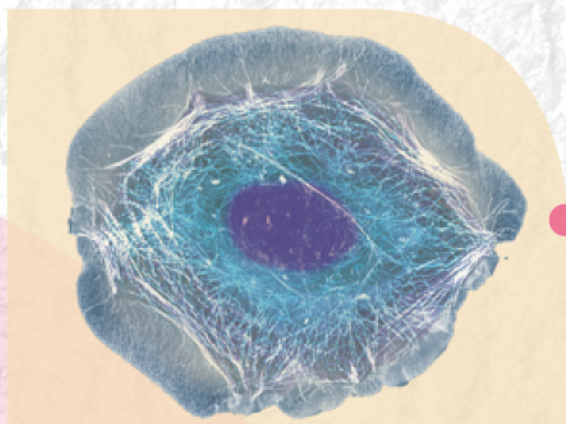
Promotes deposition of ECM components



HA promotes organized deposition of essential components of the extracellular matrix (e.g. collagen fibers, etc.), thereby contributing to remodeling and wound healing.⁶⁻³

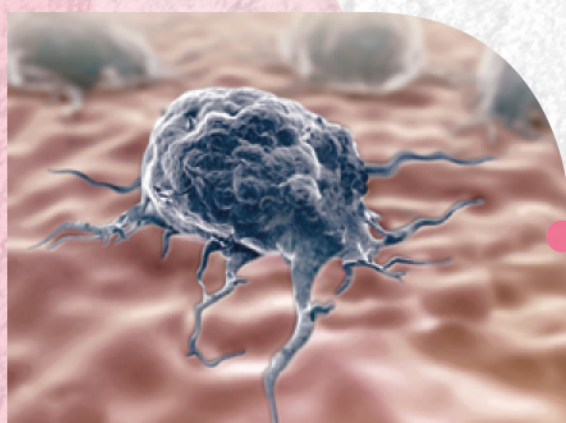
tissue repair

Migration and proliferation of keratinocytes



HA stimulates migration and proliferation of keratinocytes through interaction with CD44 receptors. Keratinocytes are responsible for restoring the epidermis after injury.³

Production of antimicrobial peptides



HA interacts with the TLR2 and TLR4 (Toll-like receptors) receptors in order to mediate the production of the antimicrobial peptides beta-defensin2 by epithelial cells⁴, thus promoting early activation of tissue repair.⁵

Hyalo4[®] Skin:

Cream containing:

- %0.2 hyaluronic acid

INDICATIONS¹: Hyalo4[®] Skin cream is indicated for the management of cutaneous irritations and lesions. In particular, it is intended to cover acute and chronic wounds (abrasions, donor sites and post-operative incisions, first and second degree burns, vascular and metabolic ulcers and pressure sores) and to provide a moist wound environment, thus protecting against abrasion, friction and desiccation.⁷

Clinically-demonstrated efficacy

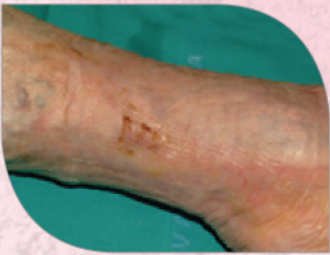
Female patient, -69years-old, with saphena bilateral venous insufficiency and very painful vascular ulcers.

RESULTS



Baseline

After aggressive surgical debridement and application of a dermal substitute Hyalo4® Skin was applied daily.



Final observation (25 days)

Wound is completely healed.

Notice the extension of tissue repair, with no evident scarring.

Key points

Hyalo4® Skin promotes proliferation and migration of cells involved in tissue repair³⁻¹

- Promotes re-epithelialization³
- Induces neoangiogenesis³
- Stimulates remodeling⁵⁻³
- Favors scarless healing³
- A wide range of formulations for the most diverse needs⁷

Hyalo4® Skin

Product description

Hyalo4® Skin Cream is a topical preparation constituted by hyaluronic acid as its principal component. Hyaluronic acid is a biological polysaccharide (glycosaminoglycan) distributed in the extracellular matrix of most tissues. Due to its hydrophilic properties it provides hydrated space around and between cells, thus facilitating the migration of cells. Hyalo4® Skin Cream is available in different forms (Hyalo4® Skin Cream, Gauze-pads, Spray and Gel).

Indications

Hyalo4® Skin Cream is indicated for the management of cutaneous irritations and lesions. In particular, it is intended to cover acute and chronic wounds (abrasions, donor sites and post-operative incisions, first and second degree burns, vascular and metabolic ulcers and pressure sores) and to provide a moist wound environment, thus protecting against abrasion, friction and desiccation.

Advantages

- Hyaluronic acid stimulates the healing processes which are faster and improved³.
- Wide ranges of formulations for the most diversified requirements¹.

Mechanism of action

Hyaluronic acid aids the migration of the fibroblasts and endothelial cells. Hyaluronic acid alters the inflammatory phase by acting as a scavenger of the free radicals and activating a negative feedback. In the remodelling phase, it reduces fibrosis and regulates the collagen secretion.

Composition

Cream, Gel, Spray. Principal component: Hyaluronic acid sodium salt at %0.2

Gauze-pads. Principal component: Hyaluronic acid sodium salt at %0.05

Product description	Format	Articles per pack
Hyalo4® Skin	15 g tube cream	1
Hyalo4® Skin	25 g tube cream	1
Hyalo4® Skin	30 g tube cream	1
Hyalo4® Skin	100 g tube cream	1
Hyalo4® Skin	30 g tube gel	1
Hyalo4® Skin	100 g tube gel	1



REFERENCES: **1.** Chen WY, Abatangelo G. Wound Repair Regen. 1999 Mar-Apr;89-79;(2)7. Functions of hyaluronan in wound repair. **2.** David-Raoudi M, et al. Differential effects of hyaluronan and its fragments on fibroblasts: relation to wound healing. Wound Repair Regen87-16:274;2008. **3.** Prosdocimi M, Bevilacqua C. Exogenous hyaluronic acid and wound healing: an updated vision. Panminerva Med35-54:129;2012. **4.** Gariboldi S, et al. Low molecular weight hyaluronic acid increases the self-defense of skin epithelium by induction of beta-defensin2 via TLR2 and TLR4. J Immunol 10-2103;181;2008. **5.** Jiang D, et al. Hyaluronan in tissue injury and repair. Annu Rev Cell Dev Biol 61-23:435;2007. **6.** Yates CC, et al. Skin wound healing and scarring: fetal wounds and regenerative restitution. Birth Defects Res C Embryo Today 33-325 (4)96 ;2012. **7.** Hyalo4® Skin. Instructions for use. Rev 11.