



RSL Revolutionary Labs is an EU-based biotech company that was created with a vision to combine natural ingredients and innovative biomaterials with state of the art formulations to offer solutions for oncology patients. RSL comes in to fill an unmet need for oncology patients, as available options for skin care management in cancer patients are still quite limited and products that are available are not effective or may contain harmful chemicals.

After years of research, expert scientific knowledge and with nature's most exquisite ingredients, RSL has created the εŪSKIN[®] line of products, to nourish and support the skin against the side effects of cancer treatment.



THE NEED

In recent years, the management of cancer as a complex disease has seen advancements through the emergence of novel chemotherapeutic agents, targeted biological therapies, and radiation protocols, which have contributed to an overall enhancement in the survival rates of oncology patients.

However, these treatments frequently result in undesirable cutaneous side effects, leading to both physical and psychological distress, and substantially impeding patients' quality of life. Therefore, the improvement of patients' quality of life has become an essential therapeutic objective in modern cancer management.



THE PROBLEM

Dermatological side effects, such as skin rash, from oncology treatments (radiotherapy, brachytherapy, immunotherapy, and chemotherapy) is probably the most common side effect that can affect patients' quality of life as well as possible termination of their treatment.

According to international literature data¹, 95% of patients with breast, skin and head cancer who undergo radiation therapy manifest radiation dermatitis (skin damage caused by radiation during radiation therapy). Therefore, the skin around the irradiated area may experience pain, erythema and even dry or wet scaling.

It is estimated that nearly 85%-90% of patients receiving radiation therapy will experience a moderate to severe skin reaction, with the most common side effect being radiation dermatitis. Several factors appear to influence the severity, onset, and duration of skin reactions.

In acute actinic dermatitis there is erythema, edema, inflammation, cell apoptosis and necrosis. Erythema is due to inflammation caused by the dilation of superficial vessels and is a result of the concentration of immune system cells and the cytokines they produce². The Radiation Therapy Oncology Group (RTOG) grading of acute radiation dermatitis describes four grades of skin reaction³, from simple erythema to skin ulceration and necrosis⁴. Measures to prevent and treat radiodermatitis include preparation and adequate hydration of the skin in the treatment area using moisturizing products.



THE SOLUTION

Regrettably, to date, the availability of specialized products catering to the unique requirements of oncology patients has been either scant or altogether absent. In response, we have developed a series of specialized products utilizing natural ingredients and innovative biomaterials to fulfill the specific needs of individuals undergoing oncological treatment and enhance their overall quality of life.





Hormone

therapy



& Biologicals



Skin inflammation (bumps, tags and rashes)

Common skin conditions

during oncological treatment



Oncology treatments can cause skin irritation and inflammation by different routes, such as damaging the rapidly dividing skin cells, therefore inducing an inflammatory response. These may lead to skin inflammation which refers to a state where the skin becomes red, swollen and itchy. It is a natural mechanism that aids in combating harmful pathogens and healing⁵.



Cracked nails - Brittleness



Oncology treatments can have an impact on the quickly dividing cells in the nail matrix, leading to various alterations in the nails. These changes often manifest as brittle, dry, and fragile nails, accompanied by cuticles that are more susceptible to cracking and splitting. Patients undergoing oncology treatments have frequently reported experiencing these nail-related issues⁶. The primary reason behind nail brittleness and cracked nails is attributed to the breakdown of the intercellular cement substance that binds the horny lamellae of the nail together. As a result, the nails tend to split in layers, often starting from the free edge. These nail alterations are among the potential side effects that patients undergoing oncology treatments may encounter⁷.



Stomatitis – Oral mucositis (oral blisters, difficulty in ingestion)



Stomatitis, a frequent adverse effect of chemotherapy, is marked by inflammation and tenderness in the mouth or the mucous membranes that line the oral cavity. Its symptoms encompass pain, redness, swelling, and the formation of mouth sores or ulcers⁸. Oral mucositis is distinguished by the presence of red and ulcerative lesions on the oral mucosa, which can cause considerable pain and may negatively impact nutrition and oral hygiene. Additionally, these lesions heighten the risk of both local and systemic infections⁹.



Atopic dermatitis (itchiness, sores)



Patients undergoing oncology treatments frequently encounter a compromised immune system, making them more susceptible to skin infections, including atopic dermatitis. Atopic dermatitis is an inflammatory skin disorder characterized by itchy skin lesions. It is a prevalent condition that exhibits eczematous lesions and intense itching, significantly impacting the patient's quality of life¹⁰.





Eczema, also known as Atopic dermatitis, is a prevalent chronic and recurrent condition marked by pruritus and a disruption of the epidermal barrier function¹¹. This skin condition leads to inflammation, irritation, and redness¹².





Hormone therapy

Immune therapy & Biologicals



Common skin conditions

during oncological treatment

Radiation therapy frequently leads to radiation dermatitis, a condition that affects a significant proportion of patients undergoing treatment, with rates reaching up to 85%. This condition can result in moderate-to-severe skin reactions, characterized by distinct changes like swelling, redness, pigmentation disturbances, and tissue necrosis¹³. Radiation dermatitis can present as acute erythema and peeling, or as chronic effects, including skin thinning, visible blood vessels (telangiectasias), and fibrosis¹⁴. Acute radiation dermatitis appears as skin lesions caused by ionizing radiation and typically emerges within days or weeks after treatment, while chronic radiation effects may manifest months or even years after radiation therapy. The impact of radiation therapy on healthy tissues is evident in cell death, particularly noticeable in regenerating tissues like the epidermis and mucosal epithelia¹⁵.

Radiation dermatitis can manifest as acute erythema and desquamation, or as chronic effects including skin atrophy, telangiectasias, and fibrosis¹. Chronic radiation can be observed months or years after the radiation therapy unlike acute radiation that can be seen days or weeks after the treatment. The effect of radiation therapy has on healthy tissues can be even seen by cell death and it is easily shown on renewing tissues like the epidermis and mucosal epithelia⁴.

Acute radiation dermatitis

It is a common adverse effect in patients receiving radiotherapy. It can be observed as a burn injury of which the severity may vary according to the treatment and factors related to the patient. The burn injury may resolve weeks after its appearance but there were cases where the reaction persisted and caused complications to the treatment¹⁶. A variation of acute radiation dermatitis may be observed, known as radiation recall which is an acute inflammatory reaction usually focused in the area of the previous radiation. Drug agents that treat cancer seem to trigger the phenomenon¹⁷.



Chemotherapy extravasation (leaking of chemo drugs in the skin & wounds)



Chemotherapy extravasation is when chemotherapy drugs leak into the surrounding tissue instead of entering the intended vein. This can lead to symptoms such as pain, swelling, redness, blistering or even ulcers in the affected area and can cause damage to the surrounding tissues¹⁸.



Malignant wounds (tumors breaking through the skin)



Malignant wounds are chronic wounds that occur when cancer cells invade and destroy healthy skin or underlying tissues. These wounds are usually ulcers that may be accompanied by pain or discharge of fluid¹⁹.



The choice of every bioactive component has been made meticulously, with a thorough consideration of scientific evidence.



Gynura procumbens

source of natural antioxidants (*phenolic compounds*) Due to its high phenolic content, it protects the skin from free radicals and environmental aggressors²².

Gynura procumbens is a medicinal plant commonly found in tropical Asia. It is known for its antibacterial, antifungal, anti photoaging and antioxidant abilities. Antioxidants are known for their protective effect on the skin as they protect it from oxidative damage. Researches have shown that due to the high percentage of phenolic compounds found in Gynura procumbens, it exhibits natural antioxidant activity²², therefore making it a powerful natural antioxidant for the skin. Previous studies have shown that UV irradiation induces degradation of collagen in the skin²³. Researches have shown that properties of Gynura procumbens inhibit the expression of the proteins responsible for the degradation of collagen induced by UV irradiation²⁴.



Collagen peptides

rich in Gly-Pro-Hyp *(amino acids)* Collagen has been shown to improve healing chronic wounds in randomised clinical trials²⁵.

Collagen is the unique, triple helix protein molecule, which forms the major part of the extracellular dermal matrix (ECM) of the dermis in the skin. Together with the glycosaminoglycans, proteoglycans, laminin, fibronectin, elastin and cellular components are responsible for fibroblast migration, survival and metabolism²⁶. Collagen type I, II and III acts as a scaffold in connective tissue that is deposited early in wound healing by the fibroblasts to help with the wound healing progress²⁷⁻²⁸. Marine derived collagen I has been previously shown to promote wound healing in wound models and clinical studies²⁹⁻³¹.



Ganoderma lucidum

excellent anti-inflammatory

(triterpenes)

Ganoderma lucidum has been shown to potentially inhibit inflammatory skin conditions and promote keratinocyte proliferation³².

Ganoderma lucidum has numerous pharmacological and therapeutic properties that make it an excellent antiallergic, antioxidant, antitumor, antiviral, and antiinflammatory ingredient³³. Terpenoids and polysaccharides in Ganoderma, are being extensively studied for their antimicrobial properties on skin as it has been shown that they act on the bacterial cytoplasmic membrane³⁴. Polysaccharides isolated from Ganoderma show an antioxidant activity and protect tissues from reactive oxygen species toxicity³⁵. In addition, extracts of Ganoderma show anti inflammatory activity as it has been shown to suppress cytokines responsible for inflammation such as IL-6³⁶.



Aloe vera

rich in vitamins and minerals (glucomannan)

Glucomannan has demonstrated anti-inflammatory, tissue regeneration acceleration, and antibacterial properties³⁷.

Aloe vera has been used for centuries to treat skin injuries such as burns and eczemas because of its anti-inflammatory³⁸, antimicrobial³⁹ and wound healing properties⁴⁰. The active ingredients found in the aloe vera leaf extract consist of tocopherols, organic acids, fatty acids, sugars - mannan, and phenolic compounds. Even though polysaccharides are the main constituent of aloe vera it was shown that the biological activity of aloe vera are a result of a synergistic action of a variety of compounds³⁷. In a clinical study performed on 60 Head and Neck cancer patients undergoing radiotherapy as a treatment it was shown that aloe vera delayed radiation induced dermatitis⁴¹.



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Cannabis seed oil

rich in cannabidiol (cannabidiol)

A valuable source of biologically active substances that reduce oxidative stress, inhibit skin aging processes and positively affect the viability of skin cells⁴².

Cannabinoids and endocannabinoids are pharmacologically active ingredients produced by cannabis sativa. Humans have an endocannabinoid system that is used to regulate several processes i.e. the production of proteins⁴³. Over 200 terpenoids are identified in C.sativa, and those are associated with medicinal properties such as antimicrobial, antioxidant and anti-inflamatory amongst other⁴⁴.Cannabis sativa seed extract impedes the mediators of inflammation that occur during wound healing⁴⁵. Cannabis sativa extract, also known as hemp extract is a valuable source of biologically active substances that reduce oxidative stress, inhibit skin aging processes and positively affect the viability of skin cells⁴².

Grapeseed oil

excellent antioxidant properties (gallic acid)

Its antioxidant properties may be beneficial to protect the skin against radiation-induced free radicals⁴⁶.

Grapeseed extract has a high composition of proanthocyanidins that have the ability to trigger the release of vascular endothelial growth factor and therefore a topical application can cause wound contraction and closure faster than the normal rate. The effect on skin lesions of the topical application of the grape seed extract increased cell density and the deposition of connective tissue at the wound⁴⁷. Grape seed extract has proven anti-inflammatory and antimicrobial properties are due to high content of polyphenols, proanthocyanidins and resveratrol⁴⁸. Polyphenolic compounds found in grape seed extract, specifically catechin and epicatechin, it is due to those compounds that the cell viability was increased and as a result protected the cells from UVA damage⁴⁹.

Panthenol

excellent moisturising abilities (pantothenic acid)

Panthenol, improves hydration in the upper layers of the skin and prevents transepidermal water loss⁵⁰.

Panthenol is very important in normal epithelial function. When applied topically, it is readily absorbed and rapidly converted enzymatically to pantothenic acid, a constituent of coenzyme A3. It acts as a moisturiser by improving hydration in the upper layers of the skin and preventing transepidermal water loss but it has also been shown to be involved in wound healing⁵⁰. As a result, panthenol has been extensively studied both for its skin moisturising/restoring abilities and for its beneficial effects in wound healing. A research group studying the effect of dexpanthenol on the skin barrier, has shown that treatment for 7 days has improved epidermal hydration and reduced transepidermal water loss⁵¹. Another clinical study focused on the efficacy of dexpanthenol in protecting skin against irritation. Their results suggest that dexpanthenol is able to protect the skin by preserving the hydration levels in the epidermis even in the presence of an irritant agent⁵². The effectiveness of dexpanthenol in wound healing has been shown by different studies, one of which focused on the topical application of dexpanthenol in in vivo models of minor skin trauma, which showed a significantly faster wound healing⁵³. Stettler et al, researched the effects of dexpanthenol in an anti scar gel in hypertrophic scars. Their results show that after 8 weeks the scars were significantly less vascularized, less pigmented, softer, thinner, flattened and more elastic⁵⁴.



Sweet orange peel oil

excellent tissue-repair properties (perillyl alcohol – POH) POH demonstrates significant anti-inflammatory effects in dermal inflammation and wound healing experiments⁵⁵.

Orange peel extracts main constituent is polymethoxyflavonoid (PMFs) which have shown to protect the skin against UV damage⁵⁶. Citrus peel extracts have been used for their antioxidant and anti-inflammatory properties⁵⁷. Dominant compound of the peel oil are monoterpene hydrocarbons specifically limonene⁵⁸. D-Limonene and its metabolite perillyl alcohol contributes significantly as an anti-inflammatory agent in murine dermal inflammation and wound-healing. D-limonene, amongst other activities assisting the wound healing process, decreases the cytokine production contributing to the reconstruction of the epidermal barrier⁵⁵.







The choice of every bioactive component has been made meticulously, with a thorough consideration of scientific evidence.



Hyaluronic acid

excellent moisturising abilities (*hyaluronic acid*) Hyaluronic acid has been shown to reduce the surface area of the wound by 70%⁵⁹.

Hyaluronic acid is a natural polysaccharide and a key component of the extracellular matrix and is known to be involved in several mechanisms of the wound healing process such as decreasing inflammation, regulating tissue remodeling and enhancing angiogenesis⁶⁰. Skin aging, both intrinsic and extrinsic (ie. by UV irradiation), is associated with loss of moisture in the skin. One key molecule responsible for maintaining the moisture in the skin is hyaluronic acid with its unique ability to bind and retain water molecules⁶¹, thus it plays a vital role in skin aging. In a clinical trial studying wound healing in acute wounds of patients, it has been shown that hyaluronic acid was able to help decrease the wound by 70% in size, 10 days after application⁵⁹. In 2012, a systematic review was published, which collected clinical trials on the usefulness of hyaluronic acid derivatives in the treatment of a wide variety of wounds: burns (including radiodermatitis), superficial surgical wounds (dermabrasion) and chronic wounds (venous ulcers and diabetic foot)⁶².

Balsam oil

rich in naphthoquinones (naphthoquinones)

Naphthoquinones found in Balsam oil possess remarkable wound healing and anti-inflammatory activities⁶³.

Hypericum perforatum has been used both orally and topically for healing wounds and burns probably due to its antioxidant, antimicrobial and anti inflammatory properties⁶⁵. A clinical study showed that topical application of the extract on cesarean sections promoted healing and epithelial reconstruction⁶⁵. Another clinical study has shown an increase in hydration and a reduction in transepidermal water loss when hypericum was applied topically in comparison to the control group⁶⁵. In addition, St John's Wort has been shown to aid with wound healing of burn wounds. When applied topically, it aided acceleration of healing of second and third degree burn wounds 3 times faster than conventional methods⁶⁶.



Calendula officinalis

excellent anti-inflammatory (xanthophyll)

Xanthophyll has a role in treating minor inflammation of the skin and assisting the healing process of minor wounds⁶⁷.

Calendula officinalis flower extract has been widely used for the treatment of minor wounds, and it has been approved by the European Medicine Agency to be used in products that aim to reduce inflammation in the skin. In a study using scratch assays, it has been shown that Calendula had an effect on the inflammatory phase of wound healing by activating a pathway that increases IL-8 in the keratinocytes thus increasing wound closure⁶⁷. Preethi et al, has found that when the calendula extract was used in an in vivo wound healing model, it led to reduced proinflammatory markers and reduced edema⁶⁸. Calendula also improves distensibility, firmness and viscoelasticity in the skin⁶⁹.



Shea butter

excellent source of fatty acids (fatty acids)

Fatty acids are the main component of shea butter that play a role in its antioxidant and anti-inflammatory properties $^{\rm 70-72}$.

Shea butter is the solid fat extracted from mature fruits of the sheu tree(Vitellaria paradoxa). 90% of its constituents are triglycerdes and 10% non-triglycerides such as oil soluble tocopherols, triterpenes, phenols, allantoin, polyphenols and karitene. Shea butter is a rich source of fatty acid, more common to found stearic, oleic, palmitic, linoleic and arachidic⁷³.Topical use of Shea butter has shown anti-aging and anti-inflammatory properties⁷⁴. The anti-inflammatory properties of shea butter were proved by the inhibition of iNOS, COX-2 and cytokines⁷⁵.



The choice of every bioactive component has been made meticulously, with a thorough consideration of scientific evidence.



Cucumber extract

excellent antioxidant properties (vitamin C)

Rich in Vitamin C, it stimulates collagen synthesis and assists in antioxidant protection and photodamage⁷⁶.

Cucumis sativus extract has shown antioxidant and analgesic activity. Compounds found in the extract responsible for those activities are flavonoids and tannins⁷⁷. The extract is also rich in vitamin C which stimulates collagen synthesis and assists in antioxidant protection and photodamage⁷⁸⁻⁷⁹. Cucumber is used not only for its soothing effects on skin irritation but also reduces swelling. Cucumis sativus extracts have shown pharmacological activities such as antioxidant, antiwrinkle and antiaging and antimicrobial. Lactic acid an ingredient found in cucumber juice is used for dry skin, ichthyosis⁷⁶. A topical skin care cream produced with cucumber extract was proven to increase trans-epidermal water loss and acts as a whitening agent⁸⁰.



Chamomile extract

rich in bisaboloids (*a-bisabolol molecule*) Has a role in faster reepithelialisation and wound-breaking strength⁸¹.

Chamomile, a medicinal plant, contains levomenol, bisaboloids, chamazulene, and flavonoids, which are responsible for its anti inflammatory and antimicrobial properties⁸². It has been shown that when it is applied topically as a gel, it delays the onset of radiation dermatitis in a study conducted with head and neck cancer patients undergoing radiotherapy⁸³. Another study by Nayak et al has shown that when chamomile extract was applied topically on the wound, it was able to reepithelialise faster and had a significantly higher wound-breaking strength in comparison to the control group⁸¹.



Lavender essential oil

rich in monoterpenes (linalool molecule)

Due to its high monoterpene content, it has anti-bacterial and anti-fungal properties⁸⁴.

Lavender oil owes its anti-bacterial and anti-fungal activity to its main components monoterpenes such as linalool and linalyl acetate⁸⁴. It was shown in a study performed on rat models that topical application of lavender oil increased collagen synthesis by fibroblasts⁸⁵. It was proven that lavender essential oil is an inhibitor of the synthesis of 4 pro-inflammatory cytokines characterizing the oil for anti-inflammatory treatment⁸⁶.

INTENSIVE CREAM

The **EUSKIN®** Intensive Cream is specifically formulated to aid in the recovery of skin damaged by radiotherapy, chemotherapy, or other irritants. It is composed of a range of natural and restoring ingredients including Ganoderma lucidum, Hyaluronic acid and Panthenol.



Ganoderma lucidum extract has numerous pharmacological and therapeutic properties that make it an excellent antiallergic, antioxidant, antiviral, and anti-inflammatory ingredient³⁴.



Hyaluronic acid is known to be involved in wound healing processes such as decreasing inflammation, regulating tissue remodelling and enhancing angiogenesis⁶⁰. It is also important for maintaining the moisture in the skin with its unique ability to bind and retain water molecules⁶¹.



Panthenol functions as a moisturizer, inhibits trans-epidermal water loss^{50,51} and accelerates the process of wound healing⁵³.



APPLICABLE SKIN CONDITIONS

Papulopustular eruption, Skin rash, Radiation dermatitis, Irritative dermatitis, Pruritus, Dry skin, Erythema, Wound healing complications, Urticaria rash, Atopic dermatitis, Eczema, Skin atrophy, Skin wounds, Cheilitis.

RECOVERY OIL

εΰSKIN® Recovery Oil's non greasy formula contains Gynura procumbens, Cannabis sativa and Grapeseed to assist with the recovery of sensitive and irritated skin areas.



Gynura procumbens was found to inhibit the expression of the proteins responsible for the degradation of collagen²⁴. Due to its high percentage of phenolic compounds, it exhibits natural antioxidant activity²².



Cannabis seed oil has been found to prevent skin inflammation that occurs during the process of wound healing⁴⁵.



Grapeseed oil has anti-inflammatory and antimicrobial properties due to the high content of polyphenols, proanthocyanidins and resveratrol. Topical application in skin lesions increases cell density and deposition of connective tissue at the wound⁴⁷.



APPLICABLE SKIN CONDITIONS

Hand-foot syndrome (HFS), Palmar-plantar erythrodysesthesia (PPE) - Bullous acral erythema, Xerosis, Dry skin, Acquired ichthyosis, Wound healing complications, Sebaceous hyperplasia, Eczema, Skin ulcers, Petechiae, Skin wounds.



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SOOTHING BODY LOTION

The **£USKIN®** Soothing Body Lotion is a nutrient-rich product that features Collagen peptides, Balsam oil, and Hyaluronic acid. It also contains Aloe vera and Gynura procumbens, known for their anti-inflammatory⁴¹, antimicrobial³⁹, and antioxidant²⁹ characteristics.



Collagen peptides have been previously demostrated to promote healing in wound skin models and clinical studies²⁹⁻³¹.



Balsam oil has been utilized for the treatment of wounds and burns both orally and topically, likely due to its antioxidant, antimicrobial, and anti-inflammatory properties⁶⁴. Moreover, the application of the extract on caesarean sections resulted in faster healing and epithelial reconstruction⁸⁷.



Hyaluronic acid has been recognized for its role in wound healing, which includes the ability to reduce inflammation, regulate tissue remodelling, and promote angiogenesis⁶⁰. It also plays a crucial part in preserving the skin's hydration levels by its distinctive capacity to attach and hold onto water molecules⁶¹.



APPLICABLE SKIN CONDITIONS

Radiation dermatitis, Irritative dermatitis, Pruritus, Dry skin, Xerosis, Acquired ichthyosis, Wound healing complications, Urticaria rash, Atopic dermatitis, Skin ulcers, Skin atrophy, Skin wounds.

INTENSIVE GEL

The **£USKIN® Intensive Gel** is a formulation that is also suitable for use in mucosal areas. It contains Aloe vera, Cucumber, and Chamomile extracts, which imbue the gel with anti-inflammatory^{38,82}, antimicrobial^{76,82}, analgesic⁷⁷ and refreshing capabilities.



Aloe vera has been used for centuries to treat skin injuries such as burns and eczema owing to its anti-inflammatory³⁸, antimicrobial⁴⁰ and wound healing properties⁸⁸. In a clinical study it was also revealed that Aloe vera had a delaying effect on radiation-induced dermatitis in patients who were undergoing radiotherapy.



Cucumber extract has been shown to have antioxidant and antimicrobial properties probably due to its high content of flavonoids and tannins⁷⁶⁻⁷⁷. The extract is also rich in Vitamin C, which stimulates collagen synthesis and assists in antioxidant protection and photodamage⁷⁸⁻⁷⁹.



Chamomile extract has anti-inflammatory and antimicrobial properties⁸² attributed to its constituents such as levomenol, bisaboloids, chamazulene, and flavonoids. Topical application of chamomile extract on the wound has resulted in faster reepithelialisation compared to the control group⁸¹ in a clinical setting.





NAIL REPAIR OIL

εΰSKIN® Nail Repair Oil was developed with the purpose of enhancing the growth of healthy nails and repairing damaged ones. Its key components are Grapeseed, Lavender, and Gynura procumbens.



Grapeseed oil has anti-inflammatory and antimicrobial properties due to the high content of polyphenols, proanthocyanidins and resveratrol. Topical application on skin lesions increases cell density and deposition of connective tissue at the wound⁴⁷.



Lavender oil owes its anti-bacterial and anti-fungal activity to its main components, which are monoterpenes such as linalool and linalyl acetate⁸⁴. It was previously shown that topical application of lavender oil increased collagen synthesis by fibroblasts⁸⁵.



Gynura procumbens properties were found to inhibit the expression of the proteins responsible for the degradation of collagen²⁴. Due to its high percentage of phenolic compounds, it exhibits natural antioxidant activity²².



APPLICABLE SKIN CONDITIONS Paronychia, Onycholysis, Cracked nails, Brittleness, Dystrophic nails.

LIP CARE

The **£USKIN®** Lip Care is a mild fusion of rejuvenating and moisturizing constituents, specially created to soothe dryness and cracks in the lips. Its principal elements consist of Sweet orange peel oil, Calendula, and Shea butter.



Sweet orange peel oil is mainly composed of monoterpene hydrocarbons, specifically limonene⁵⁸. D-Limonene contributes as an anti-inflammatory agent in dermal inflammation and wound-healing⁵⁵.



Calendula officinalis flower extract has been approved by the EMA to be used in products that aim to reduce inflammation in the skin. It has been shown it can impact the inflammatory phase of wound healing by activating a pathway that increases IL-8 in the keratinocytes, thus increasing wound closure⁶⁷.



Shea butter is a rich source of fatty acids, such as stearic, oleic, palmitic, linoleic and arachidic⁷³. The topical use of Shea butter has shown anti-aging and anti-inflammatory properties⁷⁴.



APPLICABLE SKIN CONDITIONS Stomatitis, Oral mucositis, Perioral dermatitis, Cheilitis.



GENTLE WASH

The <code>ɛ̃ŪSKIN® Gentle Wash</code> gently cleanses and maintains the skin's pH balance. Its key ingredients include Cucumber extract, Chamomile extract and Aloe Vera.



Cucumber extract has been shown to have antioxidant and antimicrobial properties probably due to its high content of flavonoids and tannins⁷⁶⁻⁷⁷. The extract is also rich in Vitamin C, which stimulates collagen synthesis and assists in antioxidant protection and photodamage⁷⁸⁻⁷⁹.



Chamomile extract contains levomenol, bisaboloids, chamazulene, and flavonoids, which are responsible for its anti-inflammatory and antimicrobial properties⁸². When applied topically on the wound, it was able to reepithelialise the wound faster in comparison to the control group⁸¹.



Aloe vera has been used for centuries to treat skin injuries such as burns and eczemas because of its anti-inflammatory³⁸, antimicrobial⁴⁰ and wound healing properties⁸⁸. In a clinical study, with patients undergoing radiotherapy, it was shown that aloe vera delayed radiation induced dermatitis⁴¹.

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APPLICABLE SKIN CONDITIONS

Papulopustular eruption, Skin rash, Radiation dermatitis - Irritative dermatitis, Hand-foot syndrome (HFS), Palmar-plantar erythrodysesthesia (PPE) - Bullous acral erythema, Pruritus, Xerosis, Dry skin, Acquired ichthyosis, Erythema, Wound healing complications, Sebaceous hyperplasia.



References

- Gosselin, T. K. Schneider, S.M. Plambeck, M.A. Rowe, K. (2010). A prospective randomized, placebo-controlled skin care study in women diagnosed with breast cancer undergoing radiation therapy. Oncology nursing forum, 37(5), 619–626.
- 2. Hymes, S. R. Strom, E. A. Fife, C. (2006). Radiation dermatitis: clinical presentation, pathophysiology, and treatment 2006. Journal of the American Academy of Dermatology, 54(1), 28–46.
- 3. Huang, C. J. et al (2015). RTOG, CTCAE and WHO criteria for acute radiation dermatitis correlate with cutaneous blood flow measurements. Breast (Edinburgh, Scotland), 24(3), 230–236.
- 4. Kole, A. J., Kole, L., & Moran, M. S. (2017). Acute radiation dermatitis in breast cancer patients: challenges and solutions. Breast cancer (Dove Medical Press), 9, 313–323
- 5. Alley, E., Green, R., Schuchter, L., (2002). 'Cutaneous toxicities of cancer therapy', Current opinion in oncology, 14(2), 212–216.
- 6. Mittal, S., Khunger, N., Kataria, S.P., (2022). 'Nail Changes with Chemotherapeutic Agents and Targeted Therapies' Indian Dermatol Online J. 24;13(1):13-22.
- 7. Colombo, E,V Gerber, F, Bronhofer, M, Floersheim G,L (1990). 'Treatment of brittle fingernails and onychoschizia with biotin: Scanning electron microscopy' Journal of the American Academy of Dermatology. 23(6) p 1127-32.
- 8. Wojtaszek C. (2000). 'Management of chemotherapy-induced stomatitis', Clinical journal of oncology nursing, 4(6), 263–270.
- 9. Lalla RV, Sonis TS, Peterson ED, (2008) 'Management of oral mucositis in patients with cancer' Dent Clin North Am. 2008 January; 52(1): 61-viii.
- 10. Langan, S., M., Irvine A., D. Weidinger, S. (2020). 'Atopic dermatitis', *Lancet*, 396(10523), 758.
- 11. Sohn A, Frankel, A, Patel R, V, Goldenberg G, (2011), 'Eczema' Mt Sinai J Med 78(5) p730-9.
- 12. Nemeth, V. Justin, E. (2021), 'Eczema' StatPearls[Internet] Treasure Island (FL) : StatPearls Publishing.
- 13. Rosenthal, A. Israilevic, R. Moy, R. (2019). 'Management of acute radiation dermatitis: A review of the literature and proposal for treatment algorithm', Journal of the American Academy of Dermatology, 81(2), 558–567.
- 14. Leventhal, J, Young, M, R, (2017). 'Radiation Dermatitis: Recognition, Prevention, and Management', Oncology (Williston Park), 15;31(12):885-7,894-9.
- 15. Benomar, S. Boutayeb, S. Lalya, I. Errihani, H. Hassam, B. El Gueddari B, (2010). 'Treatment and prevention of acute radiation dermatitis', Cancer/Radiotherapy, 14(3), p213-216.
- 16. Seite, S. Bensadoun, R. J. Mazer, J. M., (2017). 'Prevention and treatment of acute and chronic radiodermatitis.' Breast Cancer (Dove Med Press). 2(9), p551-557.
- 17. Burris, H, A, 3rd, Hurtig, J, (2010). 'Radiation recall with anticancer agents.' The oncologist. 15(11), p1227-37.
- 18. Jackson-Rose, et al. (2017). 'Chemotherapy Extravasation: Establishing a National Benchmark for Incidence Among Cancer Centers', Clinical journal of oncology nursing, 21(4), 438-445.
- Tsichlakidou A., Govina O., Vasilopoulos, G., Kavga A., Vastardi M., Kalemikerakis I., (2019). 'Intervention for symptom management in patients with malignant fungating wounds a systematic review', Journal of B.U.ON.: official journal of the Balkan Union of Oncology, 24(3), 1301–1308.
- 20. Yosipovitch, G. Jordan D. R., Takashi H, (2018) 'Itch: From mechanism to (novel) therapeutic approaches' Clinical reviews in allergy and immunology, 142(5), p1375-1390.
- 21. Han, L Xinzhong Dong (2014) 'Itch mechanisms and circuits' Annu Rev Biophys, 6(43): 331-335.
- 22. Rosidah, Y. M, Mun Yam, Amirin Sadikun & Mohd. Asmawi. (2008). 'Antioxidant potential of Gynura procumbens', Pharmaceutical Biology, 46(9), 616–625.
- 23. Rittié, L., & Fisher, G. J. (2002). UV-light-induced signal cascades and skin aging. Ageing research reviews, 1(4), 705-720.
- Kim, J., Lee, C. W., Kim, E. K., Lee, S. J., Park, N. H., Kim, H. S., Kim, H. K., Char, K., Jang, Y. P., & Kim, J. W. (2011). Inhibition effect of Gynura procumbens extract on UV-B-induced matrix metalloproteinase expression in human dermal fibroblasts. *Journal of ethnopharmacology*, 137(1), 427–433.
- 25. Yonath, A., & Traub, W. (1969). 'Polymers of tripeptides as collagen models. IV. Structure analysis of poly(L-proly-glycyl-L-proline)', Journal of molecular biology, 43(3), 461–477.
- 26. Tracy, L. E., Minasian, R. A., & Caterson, E. J. (2016). Extracellular Matrix and Dermal Fibroblast Function in the Healing Wound. Advances in wound care, 5(3), 119–136.
- 27. Desmouliere A, Redard M, Darby I, Gabbiani G (1995). Apoptosis mediates the decrease in cellularity during transition between granulation tissue and scar. Am J Pathol 146: 56–66.
- 28. Berry DP, Harding KG, Stanton MR, et al (1998). Human wound contraction: collagen organisation, fibroblasts and myofibroblasts. Plast Reconstr Surg 102: 124–31
- Baldursson B.T., Kjartansson H., Konrádsdóttir F., Gudnason P., Sigurjonsson G.F., Lund S.H. Healing rate and autoimmune safety of full-thickness wounds treated with fish skin acellular dermal matrix versus porcine small-intestine submucosa: A noninferiority study. Int. J. Low. Extrem. Wounds. 2015;14:37–43.
- Badois N., Bauër P., Cheron M., Hoffmann C., Nicodeme M., Choussy O., Lesnik M., Poitrine F.C., Fromantin I. Acellular fish skin matrix on thin-skin graft donor sites: A preliminary study. J. Wound Care. 2019; 28:624–628.
- 31. Woodrow T., Chant T., Chant H. Treatment of diabetic foot wounds with acellular fish skin graft rich in omega-3: A prospective evaluation. J. Wound Care. 2019; 28:76–80.
- Abate, M., Pepe, G., Randino, R., Pisanti, S., Basilicata, M.G., Covelli, V., Bifulco, M., Cabri, W., D'Ursi, A.M., Campiglia, P., Rodriguez, M., (2020). 'Ganoderma lucidum Ethanol Extracts Enhance Re-Epithelialization and Prevent Keratinocytes from Free-Radical Injury' Pharmaceuticals (Basel, Switzerland), 13(9), 224.
- 33. Quereshi S., Pandey A. K., Sandhu S. S. (2010). Evaluation of antibacterial activity of different Ganoderma lucidum extracts. J. Scientific Research Vol 3(1).
- 34. Cör Andrejč, D., Knez, Ž., & Knez Marevci, M. (2022). Antioxidant, antibacterial, antitumor, antifungal, antiviral, anti-inflammatory, and nevro-protective activity of Ganoderma lucidum: An overview. Frontiers in pharmacology, 13, 934982.
- 35. Jeong Y. U., Park Y. J. (2020). Ergosterol peroxide from the medicinal mushroom ganoderma lucidum inhibits differentiation and lipid accumulation of 3T3-L1 adipocytes. Int. J. Mol. Sci. 21, 460.
- 36. Dudhgaonkar S., Thyagarajan A., Sliva D. (2009). Suppression of the inflammatory response by triterpenes isolated from the mushroom Ganoderma lucidum. Int. Immunopharmacol. 9,
- 37. Rahman, S., Princeton, C., Bhattarai, N., (2017), 'Aloe Vera for Tissue Engineering Applications', J Funct Biomater., 8(1):6
- 38. Vasquez, B. Avila, G. Segura, D. Escalante, B.n(1996). 'Anti-inflammatory activity of extracts from Aloe vera gel.' Journal of Ethnopharmacology, 55(1) p 69-75.
- Habeeb, F. Shakir, E. Bradbury, F. Cameron, P. Travati, R.M. Drummond, A.J. Gray, A. Ferro, V.A. (2007). 'Screening methods used to determine the anti-microbial properties of aloe vera inner gel'. Methods 42 (4), p315-320.
- 40. Sanchez, M. González-Burgos, E. Iglesias, I. Gómez-Serranillos, M.P. (2020). 'Pharmacological update properties of Aloe vera and its major active constituents.' Molecules, 25(6) p 1324.
- 41. Rao, S. Hegde, S.K. Baliga-Rao, M.P. Palatty, P.L. Thomas, G. Baliga, M.S. (2017). 'An Aloe Vera-Based cosmeceutical cream delays and mitigates ionizing radiation-induced dermatitis in head and neck cancer patients undergoing curative radiotherapy: A clinical study' *Medicines* 4, 44.
- 42. Zagorska-Dziok, M., Bujak, T., Ziemlewska, A., Niziol-Lukaszewska, Z. (2021). 'Positive effect of Cannabis sativa L, Herb extracts on skin cells and assessment of cannabinoid-based hydrogel properties.' Molecules, 26(4), 802.
- 43. Martins, A, M. Gomes, A, L. Boas, I, V. Marto, J. Ribeiro H, M. (2022), 'Cannabis-based products for the treatment of skin inflammatory diseases: a timely review'. Pharmaceuticals (Basel) 15(2):210.
- 44. Nuutinen T. (2018) 'Medicinal properties of terpenes found in Cannabis sativa and Humulus lupulus.' Eur. J. Med. Chem. 157:198-228.
- Sangiovani, E. Fumagalli, M. Pacchetti, B. Piazza, S. Magnavacca, A. Khalilpour, S. Melzi, G. Martinelli, G. Dell'Agli, M. (2019) 'Cannabis sativa L. extract and cannabidiol inhibit in vitro mediators of skin inflammation and wound injury' *Phytother.* Res 33(8) 2083-2093.
- 46. Yilmaz, Y., & Toledo, R. T. (2004). Major flavonoids in grape seeds and skins: antioxidant capacity of catechin, epicatechin, and gallic acid. Journal of agricultural and food chemistry, 52(2), 255–260.
- 47. Hemmati, A, A. Foroozan, M. Houshmand, G. Moosavi, Z.B. Bahadoram, M. Maram, N.S. (2015). 'The topical effect of grape seed extract 2% cream on surgery wound healing'. *Glob J Health Sci*, 7(3):52-58.
- 48. Sochorova, L, et al (2020). 'Health effects of grape seed and skin extracts and their influence on biochemical markers'. Molecules 25(22):5311.
- 49. Yarovaya L, Waranuch N, Wisutiprot W, Khunkitti W, (2021). 'Effect of grape seed extract on skin fibroblasts exposed to UVA light and its photostability in sunscreen formulation.' J Cosmet Dermatol. 20(4):1271-1282.
- 50. Proksch, E., de Bony, R., Trapp, S., & Boudon, S. (2017). Topical use of dexpanthenol: a 70th anniversary article. The Journal of dermatological treatment, 28(8), 766–773.
- 51. Gehring, W., & Gloor, M. (2000). Effect of topically applied dexpanthenol on epidermal barrier function and stratum corneum hydration. Results of a human in vivo study. Arzneimittel-Forschung, 50(7), 659–663.
- Goujon C, Alleaume B, de Bony R, Girard P. Randomized single-blind pilot comparison study of the efficacy and tolerability of Bepanthen® Ointment in subjects with bilateral dryness of the hands. Réal Thérap Dermatol. 1997;66:47–33.
- 53. Weiser H, Erlemann GA. Beschleunigte Heilung oberflächlicher Wunden durch Panthenol und Zinkoxid. Parfüm Kosm 1987; 68: 425-8.
- 54. Stettler H, Kurka P, Kalentyeva J, et al. Clinical innovation: treatment with an anti-scar gel and massage ball improves physical parameters of hypertrophic scars. Wounds Int. 2016; 7:18–23.
- 55. D'Alessio, P.A. et al. (2014). 'Skin repair properties of d-limonene and perillyl alcohol in murine models'. Anti-infl ammatory & Anti-Allergy agents in medicinal Chemistry, 13(1), 29-35.
- 56. Yoshizaki N, Fujii T, Masaki H, Okubo T, Shimada K, Hashizume R, (2014), 'Orange peel extract, containing high levels of polymehtoxyflavonoid, suppressed UVB-induced COX-2 expression and PGE2 production in HaCaT cells through PPAR-γ activation.' *Exp Dermatol* 1:18-22.

- Hsouna A, B, Gargouri M, Dhifi W, Saad B,R, Sayahi N, Mnif W, Saibi W, (2019), 'Potential anti-inflammatory and antioxidant effects of Citrus aurantium essential oil against carbon tetrachloridemediated hepatotoxicity: A biochemical molecular and histopathological changes in adult rats.' Environmental Toxicology 34(4):388-400.
- 58. Sarrou E, Chatzopoulou P, Dimassi-Theriou K, Therios I, (2013), 'Volatile constituents and antioxidant activity of peel, flowers and leaf oils of Citrus aurantium L. Growing in Greece.' Molecules 18(9):10639-47.
- 59. Voinchet, V. Vasseur, P. Kern, J. (2006). 'Efficacy and safety of hyaluronic acid in the management of acute wounds', American journal of clinical dermatology, 7(6), 353–357.
- Cortes, H., Caballero-Florán, I. H., Mendoza-Muñoz, N., Córdova-Villanueva, E. N., Escutia-Guadarrama, L., Figueroa-González, G., Reyes-Hernández, O. D., González-Del Carmen, M., VarelaCardoso, M., Magaña, J. J., Florán, B., Del Prado-Audelo, M. L., & Leyva-Gómez, G. (2020). Hyaluronic acid in wound dressings. Cellular and molecular biology (Noisy-le-Grand, France), 66(4), 191–198.
- 61. Papakonstantinou, E., Roth, M., & Karakiulakis, G. (2012). Hyaluronic acid: A key molecule in skin aging. Dermato-endocrinology, 4(3), 253–258.
- 62. Voigt J, Driver VR. (2012) Hyaluronic acid derivatives and their healing effect on burns, epithelial surgical wounds, and chronic wounds: a systematic review and meta-analysis of randomized controlled trials. Wound Repair Regen. 20(3):317-31.
- 63. Suntar, I.P, Akkol, E.K, Yilmazer, D. Baykal, T., Kirmizibekmez, H., Alper, M., Yesilada, E., (2010). 'Investigations on the in vivo wound healing potential of Hypericum perforatum L.', Journal of Ethnopharmacology, 127(2).
- 64. Wölfle, U., Seelinger, G., & Schempp, C. M. (2014). Topical application of St. John's wort (Hypericum perforatum). Planta medica, 80(2-3), 109–120.
- 65. Heinrich U, Tronnier H. Wirksamkeit und Verträglichkeit eines Johanniskraut-Extraktes zur Pflege der atopischen Haut. Kosmetische Med 2003; 124: 133–136.
- 66. Saljic J. Ointment for the treatment of burns. Ger Offen 1975; 2: 406-452.
- 67. Cristoph, N, Junghanns, S, Hartmann, A, Murillo, R, Ganzera, M, Merfort, I,(2017). 'In vitro studies to evaluate the wound healing properties of Calendula officinalis extracts,' *Journal of Ethnopharmacology*, 196(94-103).
- 68. Preethi, K.C.; Kuttan, G.; Kuttan, R. (2009) Anti-inflammatory activity of flower extract of Calendula officinalis Linn. and its possible mechanism of action. Indian J. Exp. Biol. 47, 113–120.
- 69. Akhtar, N.; Zaman, S.U.; Khan, B.A.; Amir, M.N. (2011). Ebrahimzadeh, M.A. Calendula extract: Effects on mechanical parameters of human skin. Acta Pol. Pharm. 68, 693–701.
- 70. Badifu, G.I.O (1989). 'Lipid composition of Nigerian Butyrospermum paradoxum kernel', Journal of Oleo Science, 59, 6273-6280.
- 71. Steven M., et al. (2003). 'Phenolic constituents of shea (Vitellaria paradoxa) kernel' Journal of Agriculture and Food Chemistry, 51, 6268-6773.
- 72. Verma N. Chakrabarti R. Das R H, Gautam H K, (2012). 'Anti-inflammatory effects of Shea butter through Inhibition of Inos, Cox-2, and Cytokines via the Nf-Kb Pathway in Lps-Activated J774 Macrophage cells', Journal of complementary and integrative medicine, 9(1).
- 73. Ferreira M, S, Magalhaes M, C, Oliveira, R, Sousa-Lobo J, M, Almeida, I, F, (2021). 'Trends in the use of Botanicals in Anti-aging cosmetics'. Molecules Vol 26 Issue 12.
- 74. Malachi, O (2014) 'Effects of Topical and Dietary Use of Shea Butter on Animals.' American Journal of Life Sciences. 2 (5) p. 303-307.
- 75. Lin T, K, Zhong, L, Santiago, J, L, (2017). 'Anti-inflammatory and skin barrier effects of topical application of some plant oils.' Int J Mol. Sci 19(1):70.
- 76. Mukherjee P.K., Nema N. N., Maity N., Sarkar B.K., (2013). 'Phytochemical and therapeutic potential of cucumber', Fitoterapia, 84,227-236.
- 77. Kumar, D., Kumar, S., Singh, J., Narender, Rashmi, Vashstha, Bd., Singh, N., (2010) 'Free radical scavenging and analgesic activities of Cucumis sativus L. fruit extract! J Young Pharm 2(4):365-368.
- 78. Pullar, J,M., Carr, A,C., Vissers, M,C,M. (2017) 'The roles of Vitamin C in Skin Health' Nutrients 12;9(8):866.
- 79. Shapiro, S,S., Saliou, C,. (2001) 'Role of vitamins in skin care' Nutrition 17(10):839-844.
- 80. Akhtar N, Arshad M, Barkat A, Barkat A, K, (2011) 'Exploring cucumber extract for skin rejuvenation' African journal of biotechnology Vol.10 No.7.
- 81. Nayak, B. S., Raju, S. S., Chalapathi R., (2007). 'Wound healing activity of Matricaria recutita L. extract', *Journal of wound care*, 16(7), 298–302.
- 82. Gupta V, Mittal P, Bansal P, Khokra SL, Kaushik D. (2010) Pharmacological potential of Matricaria recutita-A review. Int J Pharm Sci Drug Res. 2:12–6.
- Ferreira EB, Ciol MA, de Meneses AG, Bontempo PSM, Hoff man JM, Reis PEDD. (2020) Chamomile gel versus urea cream to prevent acute radiation dermatitis in head and neck cancer patients: results from a preliminary clinical trial. Integr Cancer Ther. 19:1534735420962174.
- Bialon, M., Krzysko-Kupicka, T., Nowakowska-Bogdan, E., Wieczorek, P. P., (2019). 'Chemical composition of two different lavender essential oils and their effect on facial skin microbiota', Molecules, 24(18), 3270.
- Hiroko-Miyuki, M., Hiroshi K., Hirohisa K., Motokuni A., (2016). 'Wound healing potential of lavender oil by acceleration of granulation and wound contraction through induction of TGF-β in a rat model.' BMC Complement Altern Med. 16:144.
- Pandur E, Balatinacz A, Micalizzi G, Mondello L, Horvath A, Sipos K, Horvath G, (2021). 'Anti-inflammatory effect of lavender (Lavandula angustifolia Mill.) essential oil prepared during different plant phenophases on THP-1 macrophages.' BMC Complement MEd Ther. 21(1):287.
- Lavagna, S. M., Secci, D., Chimenti, P., Bonsignore, L., Ottaviani, A., Bizzarri, B.(2001). 'Efficacy of Hypericum and Calendula oils in the epithelial reconstruction of surgical wounds in childbirth with caesarean section', *Farmaco* (Societa chimica italiana : 1989), 56(5-7), 451–453.
- 88. Liang, J., Cui, L., Jiankang, L., Guan, S., Zhang, K., Jingan, L. (2021). 'Aloe vera: A Medicinal Plant Used in Skin Wound Healing', Tissue engineering. Part B, Reviews, 27(5), 455–474.
- Chu, C. N., Hu, K.C., Wu R., C., Bau D. T., (2021). 'Radiation-irritated skin and hyperpigmentation may impact the quality of life of breast cancer patients aft er whole breast radiotherapy' BMC cancer, 21(1), 330.
- 90. Mervis, J. S., & Phillips, T. J. (2019). 'Pressure ulcers: Pathophysiology, epidemiology, risk factors, and presentation' Journal of the American Academy of Dermatology, 81(4), 881–890.

