

#### P80-18 PROTEOGLYCANS PLUS



There are several parameters that significantly influence the appearance and attractiveness of the skin, such as hydration, firmness or the presence of wrinkles.

However, one of the concepts that has most influenced throughout history in the different canons of beauty is

pigmentation or skin tone.

A radiant complexion is associated with beauty. The distribution of pigment, melanin, and skin texture vary according to health status and age. Irregular coloration is an indicator of skin age, as aging is associated with the presence of spots. The difference in skin color is mainly due to the presence of melanin, a pigment that protects from external aggressions, such as UV rays.

When the body generates too much melanin to protect itself from aggressions, or simply because of aging, accumulations can be created that cause spots or changes in skin tone. This disorder is called hyperpigmentation and can affect all skin types.

**Hyperpigmentation** is the third greatest cosmetic concern, as it is one of the obvious signs of the decline of youth. For this reason, the cosmetic goal is to reduce the spots and equalize the tone of the skin.

**P80-18** illuminates the skin remarkably respecting the natural characteristics and health of the skin. In vitro tests showed that the synergistic combination of ingredients contained in **P80-18** inhibits melanin synthesis by 90%.

P80-18 is a regenerating, antioxidant and protective serum. It is a cellular energizer, Hydro-regulator, and reinforces the skin structure.

Containing a powerful polysaccharide complex that combines the action of marine glycogen and glycosaminoglycans (GAGs).

GAGs are an essential part of the extra-cellular matrix, giving support to the collagen and



elastin fibers which give elasticity and resistance to the skin. P80-18 also contains a combination of active ingredients that bring luminosity and vitality to the skin, giving it a radiant and healthy appearance immediately. Improves skin texture and elasticity, homogenizing its tone and minimizing wrinkles and fine lines.

It allows to recover the luminosity and light lost as a result of the passage of time (chronological aging), of an undue exposure to UV rays (photo-aging), as well as numerous aggressions to which our skin is exposed on a daily basis (pollution, cold, tobacco, poor diet, lack of sleep, stress, ...).

With a continuous treatment, P80-18 homogenizes the skin tone and reduces unwanted pigmentations.

Contributes to reduce wrinkles and fine lines through the synthesis of collagen and elastin

It increases the firmness, smoothness and elasticity that the skin of the face is losing as a result of the passage of years and the various aggressions to which it is exposed, since by stimulating the synthesis of collagen it contributes to regenerate the dermal matrix or tissue of support.

#### **BIOACTIVE INGREDIENTS IN P80-18**

## 1) CORUM 9515: new generation of stable vitamin C



**CORUM 9515** is a new generation of stable vitamin C derivative that provides superb whitening effects, serves to promote collage synthesis and protects DNA damage.

- ✓ Effective and stable skin lightening agent
- ✓ Balance the skin tone
- ✓ Reduce dark spot
- ✓ Prevent photoaging
- ✓ Increase collagen synthesis
- ✓ Excellent anti-oxidation properties
- √ Scavenge radical
- ✓ DNA protection

Vitamin C, or L-ascorbic acid, acts as a cofactor for collagen synthesis. It has a high regenerating ability, by its collagen synthesis stimulating activity.

Vitamin C is essential for the proline hydroxylation, therefore in the development and maintenance of collagen integrity. In addition, vitamin C inhibits the synthesis of Extracellular Matrix Metalloproteinase enzymes of, enzymes which stimulates collagen degradation in the dermis.



Vitamin C's collagen stimulating properties provides it with wound healing properties, caused by trauma, cuts, burns, or surgery. It is also suitable for the formation of new tissues.

Vitamin C belongs to the group of water soluble vitamins, and like most of them, it is not stored in the body for a long period of time, but in small quantities which are eliminated through urine. For this reason, Vitamin C daily administration is important in order to provide sufficient antioxidant protection.



Its chemical structure is similar to that of glucose (in many mammals and plants, this vitamin is synthesized from glucose and galactose). All compounds which possess the biological activity of ascorbic acid are known as Vitamin C. We should note that the only active form of Vitamin C is L-Ascorbic Acid.

As Vitamin C is a water-soluble substance, it is rapidly eliminated from the organism. Our body tends to protect vital organs, so any vitamin

deficiency is felt primarily in the skin (less vital organ), which explains the importance of its topical application.

Pure Vitamin C is very unstable and sensitive to oxidation. Vitamin C contained in **Radiance Booster** is stabilized by an ethyl group and its effectiveness has been tested clinically:

## **Clinical Efficacy Studies**

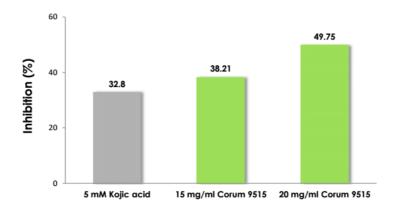
# 1- In-vitro whitening activity study - melanin assessment on human melanocytes:

An in-vitro evaluation of the Ethyl Ascorbic Acid anti-pigment ability was performed (Ethyl Ascorbic Acid is the Vitamin C form contained in VITAMIN C BRIGHTNESS SPRAY). Theophylline was incubated with melanocytes in order to increase melanin production. Subsequently, Ethyl Ascorbic Acid was added at concentrations of 15 mg/ml and 20 mg/ml respectively, and comparing with kojic acid, a known anti-pigment ingredient.

At both concentrations of Ethyl Ascorbic Acid clear depigmentation effect was observed, obtaining 49.75% whitening effect with 20mg/ml Ethyl Ascorbic Acid:



# Whitening effect (%)



## 2- In-vivo whitening efficacy

A study on the in-vivo whitening capacity of Vitamin C (as Ethyl Ascorbic Acid) for 28 days in 20 healthy Asian women aged 25 to 40 years old with skin type III was performed.

A significant improvement in the skin lightening measured by chromatography observed:



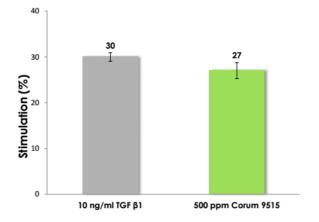


## 3- Stimulation of natural collagen synthesis:

The Stimulation of natural collagen synthesis activity was evaluated on human fibroblast culture.

After 24 hours of culture, the collagen was quantified using a Sircol Quantification Kit. Vitamin C had a similar effect on collagen synthesis that of TGF  $\beta1$  (growth factor which stimulates collagen synthesis):

## Stimulation of natural collagen synthesis:



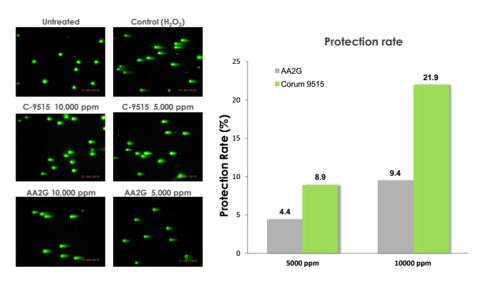


## 4- DNA protection by Comet assay in human fibroblasts:

The comet assay is a technique used to detect DNA damage and cell repair capacity. It is based on the DNA labile alkaline lysis at sites where damage has occurred.

When DNA is in good condition, it has a highly organized association with matrix proteins in the cell nucleus. When damaged, this organization is interrupted. The single DNA strands losing their compact structure and relaxes, expanding out.

Human fibroblasts were treated with Vitamin C (Ethyl Ascorbic Acid) for 24 hours, and then exposed to 100 mM H2O2. It was shown that Vitamin C at concentrations of 5000 ppm and 10,000 ppm, was able to protect DNA:



#### Vitamin C Mechanisms of action:

Vitamin C anti-aging action is exerted through several ways:

## 1. Synthesis and repair of dermal collagen



Deficiency of ascorbic acid (AA) produces significant alterations in connective tissue, since Vitamin C is essential for collagen synthesis.

Vitamin C is essential for the transformation of proline in hydroxyproline and lysine in hydroxylysine (essential constituents of collagen). Consequently Vitamin C offers stability to the extracellular matrix.

The local increase of vitamin C means therefore significantly promote collagen production; therefore improved skin elasticity and greater resistance in wall capillaries are assessed.



## 2. Antioxidant activity

Vitamin C protects cells from free radicals. From all the scientific publications regarding Vitamin C, the most interesting are those related to the photoprotective effect of ascorbic acid when topically applied.

In mouse and pig it showed that when ascorbic acid is applied before UV radiation exposure, the negative consequences it causes in the skin (erythema, histological changes, "burned cells", wrinkles ...) decreased significantly.

A study published by the Journal of Investigative Dermatology in May 1996, describes how topical application of vitamin C, vitamin E and Selenium protects rats skin cells from damage caused by exposure to UVB rays.

In parallel, the British Journal of Dermatology some years before, evidenced this protective effect of vitamin C, when used topically, on pig skin damaged by ultraviolet radiation.

The Spanish Journal of Physiology published a study showing how direct application of vitamin C protects, and thus prevents the aging in human skin cells in culture subjected to a strong oxidation stimulus with hydrogen peroxide.

One might think that its photoprotective effect was physical, that is to say, topical vitamin C behaves as a sunscreen, and however, its absorption spectrum has nothing to do with the emission of UV radiation. Later it was found that UV radiation produced a significant decrease in the levels of ascorbic acid in the skin.

All this goes to show that the UV light, after exhausting all the vitamin C present in the skin, cause an increase in free radicals, making manifest the neutralizing action of vitamin C.

## 3. Anti-inflammatory action

Vitamin C inhibits NFkB, which is responsible for the activation of a number of proinflammatory cytokines. Therefore, Vitamin C has a potential anti-inflammatory activity and can be used in conditions like acne vulgaris and rosacea. It can promote wound healing and prevent post-inflammatory hyperpigmentation.

#### 4. Vitamin C as a whitening agent

When choosing a whitening agent, it is important to differentiate between substances that are toxic to the melanocyte and substances that interrupt the key steps of melanogenesis. Vitamin C falls into the latter category of depigmenting agents. Vitamin C interacts with copper ions at the tyrosinase-active site and inhibits action of the enzyme tyrosinase, thereby decreasing the melanin formation.



## 2) PURIFIED MARINE GLUCOGEN: Energizing, restorative, anti-stress



Glycogen provides the necessary substances to restore the energy of epidermal cells and stimulate cellular metabolism.

It has a protective, regenerative and restorative function, it is a specific energizer with revitalizing, regenerating and strengthening effects.

Reinforces epidermal defense and it restorative capabilities.

Glycogen is a reserve polysaccharide, constituted from glucose molecules. It helps decrease the damage that occurs in the stressed skin, increasing oxygen consumption in epithelial cells. Glycogen is one of the main energy sources for cellular activity and metabolism. It is a source of energy reserve since it is stored for later use when necessary.

The energy obtained from glycogen is used for cell defense and restoration.

Increases oxygen consumption in cells helps fight cell stress.

#### 3) OCEOVITAL GAG: Proteoglycans

Oceovital GAG is a powerful polysaccharide complex that combines the action of marine-derived glycogen and glycosaminoglycans.

Glycogen is the energy source of skin cells. It is present in all defense and restoration processes carried out by the skin cells as an energy supply. It also improves oxygen consumption and acts as an anti-stress ingredient against external stressors such as irradiation, extreme temperatures, pollution ...

The skin's extra-cellular matrix is like a gelatinous substance in which the cells layers are floating. The glycosaminoglycans are an essential part of this extra-cellular matrix, giving support to the collagen and elastin fibers which give the elasticity and resistance to the skin. In addition, the production and maintenance of the glycosaminoglycans and extracellular matrix is essential to maintain dermal hydration, since they have a high affinity for water



molecules and are therefore very effective as natural moisturizers providing moisture to the skin leaving it hydrated and fresh.

The loss of the extra-cellular matrix glucosaminoglycans weakens the skin and results in a flaccid skin, without firmness and with great dehydration.

The main glycosaminoglycans present in this active are chondroitin sulfate and hyaluronic acid.

Oceovital GAG is a cellular energizer, Hydro-regulator, and reinforces the skin structure.

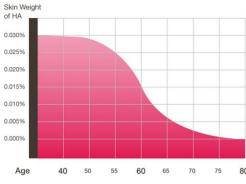
## 4) HYALURONIC ACID:



Hyaluronic acid (HA) is a polysaccharide from the type  $\beta$ - links glycosaminoglycans, having a structural function, such as chondroitin sulfates. It has the capacity to absorb more than 1000 times its volume in water.

That is why it is used in epidermis moisturizing the as

it reconstructs the fibers that hold skin tissues, giving a better shape. With a very high viscoelasticity, it is a natural component part of the skin and is essential to fight aging and wrinkles due to its high moisturizing power.



In the dermis, hyaluronic acid is the main component of the extracellular matrix (ECM). Fibroblasts are a cell type responsible for the production of collagen and elastin in the skin. ECM extracellular matrix is the space between the skin cells. This makes the skin soft, smooth and elastic.

Age 40 50 55 60 65 70 75 80 Young skin (soft and elastic) contains high amounts

of HA (Hyaluronic acid).

Hyaluronic acid contained in GOLD 24K

Hyaluronic acid contained in GOLD 24K FLASH, is of biotechnological origin, has a molecular weight of 50-110 kDa:

- ✓ Retains moisture and elasticity in the tissues (moisture retention in the extracellular matrix (ECM))
- ✓ Protects against environmental stress
- ✓ Helps to reduce the appearance of wrinkles and expression lines.



## 5) ALOE VERA: Excellent moisturizer



Aloe juice is obtained from the pulp of its fleshy leaves by physical processes. It consists of a complex mixture of more than 20 substances, including mono and polysaccharides, anthraquinones, enzymes, vitamins (A, B1, B2, B6 and B12), salicylic acid, saponins, sterols, and minerals [4]. It's

moisturizing, soothing, anti-inflammatory, anti-allergic and regenerative properties are well known.

**Fatty acids:** cholesterol, campesterol, β-sisosterol and lupeol. All have anti-inflammatory action. Lupeol also has antiseptic and analgesic properties.

It contains **phytohormones**: **auxins and gibberellins** that aid in wound healing and have anti-inflammatory action [5].

Aloe juice has been shown to **enhance the cellular structure of fibroblasts** and to **accelerate the process of collagen synthesis** [6].

It is an excellent moisturizer, so it captures atmospheric water by moisturizing in depth as it acts on the three layers of skin: epidermis, dermis, and hypodermis.

# 6) CALENDULA EXTRACT: Re-epithelizing and healing



Calendula is one of the best-considered plants for treating sensitive skin.

It is one of the most recognized plants in medicinal treatments. It is composed of flavonoids and triterpenes whose properties are beneficial to treat the different conditions of the skin.

It is ideal for the dry and scaly skin as it moisturizes, rejuvenates the skin, improving its elasticity.

Los extractos de las flores de C.officinalis muestran un amplio espectro de acciones farmacológicas, de ahí la gran importancia de los extractos de caléndula en la cosmetología moderna [1].

# Reepithelizing and wound healing activity

This is one of the most extensively used actions of calendula. It is due to the presence of polysaccharides (mucilage), flavonoids (especially quercetrin-3-O-glycoside), triterpenes



and carotenes. These substances act on the metabolism of glycoproteins and on the collagen fibers. Creams containing calendula floral extract 5% in combination with allantoin, promoted remarkable epithelization with especial intensity on the metabolism of glycoproteins and collagen fibers during tissue regeneration. More recent research suggested that the water extracts of calendula flowers, applied on skin wounds, play a role as micro-vascularization inducing agents, thus contributing to speed up healing [1]. According to ESCOP, calendula (flower) is recommended for the local treatment of skin inflammation, and as a co-adjuvant in wound-healing [2].

The calendula extract polysaccharides have concentration-dependent adhesive effects (absorption effects) on the epithelial tissue of the oropharyngeal mucosa. This action contributes to its therapeutic effects to treat oropharyngeal mucosae inflammation (www.fitoterapia.net).

For these reasons, the Calendula extract is highly recommendable to formulate cosmetic products with tissue regeneration action.

## Moisturizing activity

This activity of calendula is due to its saponin and mucilage content. These active principles have moisturizing properties (water retention and water release to the medium).

Due to its moisturizing activity, Calendula is ideal for formulating cosmetic products for the dry, irritated or delicate skin.

# **Anti-inflammatory activity**



The topical use of calendula preparations is rather extensive in cosmetics as well as in dermatology because of its **anti-inflammatory activity**. The clinical efficacy of this plant has been long demonstrated. In vivo studies using rats evidenced the anti-inflammatory activity (on inflammation induced by carrageenan and prostaglandin E1) and the inhibitory action on

leukocyte infiltration.

Thus, Calendula is highly recommended to formulate cosmetic products for sensitive and/or irritated skin.



## 7) HAMAMELIS VIRGINIANA: Decongestant and antioxidant



El extracto de hamamelis se obtiene a partir de las hojas de Hamamelis virginiana L. Hamamelis tiene propiedades medicinales sobradamente conocidas. Entre sus propiedades cosméticas se encuentran las siguientes:

## Activity on blood circulation

This activity is due to the flavonoids content of hamamelis extract. Several tests demonstrated that hamamelis flavonoids have phlebotonic, vasoprotective and capillarotropic effects, and that they

reduce blood capillaries permeability. These effects are reinforced by the hamamelis leucoanthocyanidins.

- Antimicrobial activity: Due to its content in tannins and proanthocyanidins, it has bacteriostatic and antiviral properties (Alonso, J., 2004).
- Antioxidant activity: Its content in tannins, especially hydrolyzable tannins, and flavonoids inhibit lipid peroxidation.

## 8) ASIAN CENTELLA EXTRACT: Dermoprotector



Asian centella (Hydrocotile asiatica), contains an essential oil, triterpene saponins (asiaticosides, brahmosides), alkaloids (hidrocotilina), tannins, phytosterols, vitamin K and mineral salts (magnesium, calcium and sodium). Highlights include its astringent, antiseptic, anti-inflammatory and dermoprotective virtues. The triterpene derivatives stimulate the formation of fibroblasts, cells indispensable for repairing damaged tissues. In

addition, they regulate the incorporation of collagen to the skin.

## Collagen synthesis stimulating activity

The active ingredients of the selected triterpenes of Centella asiatica have shown to have modulating properties on the development and metabolism of connective tissue.

Centella asiatica has been documented to aid wound healing in several scientific studies. One of the primary mechanisms of action of this plant appears to be the stimulation of type-1 collagen production.

Triterpenes of Centella asiatica may help to improve wound repair with a better reepithelialisation and a normalisation of perivascular connective tissue allowing an improvement of the venous wall tone and elasticity.