

# Unisooth PN-47

A complete reduction of pro-inflammatory factors  
for an instant soothing

Skin irritation is amplified and controlled by various signaling pathways. By acting directly on the causes of skin irritation, Unisooth PN-47 provides an instant soothing response:

- 1) Curative action: inhibition of one of the major signaling pathways: JAK/STAT
  - › Instant interruption of cross communication between cells;
- 2) Preventive action: control of 12 chemokines and cytokines release
  - › Diminution of the pro-inflammatory factors for a prolonged effect.

instant  
inflammation  
extinguisher

# Focus on the product

## Skin irritation is a real concern for women

Almost 3 out of 4 women claim to have an irritated skin<sup>1</sup>. Discomfort, tightness, dryness and redness of the face and body are the major signs of such irritation.

Irritation and skin sensitivity are triggered by aggressions such as shaving, waxing, peeling, sun exposure or laser treatments. Therefore, the need for instant soothing is important.

## A major pathway in skin irritation: the JAK/STAT signaling pathway

Skin irritation is sustained by a cross-talk mechanism between keratinocytes in the epidermis layer and the infiltrating immune cells. This leads to an amplification loop creating an over-reaction inside the skin.

The JAK/STAT signaling pathway has one of the most important roles in sustaining this amplification loop. By providing the cross-talk mechanism, the production of cytokines is increased.

## Unisooth PN-47: a powerful molecular switch...

Unisooth PN-47 takes advantage of a powerful combination of panthenyl triacetate and naringenin to directly act on the causes of skin irritation. Panthenyl triacetate is well known for increasing wound healing, improving skin barrier, reducing UV-induced erythema, and more generally for its anti-irritation properties. On the other hand, Naringenin, which is extracted from citrus fruits, is known to inhibit pro-inflammatory cytokines in fibroblasts<sup>2</sup> and more recently for its capacity to decrease inflammatory signaling pathways such as JAK/STAT<sup>3</sup>.

## ... for an instant soothing effect

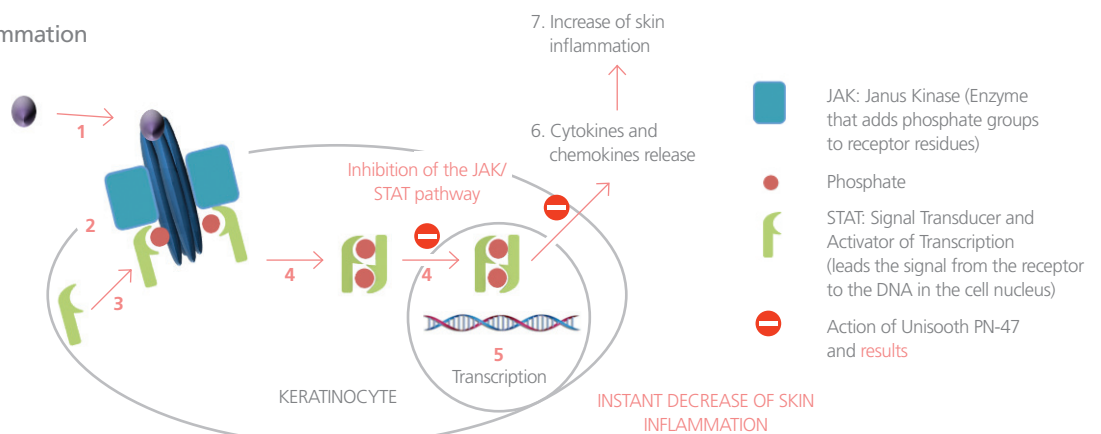
Unisooth PN-47 has a strong multi-level mechanism of action:

- Curative: it inhibits one of the major inflammatory pathway, the so called JAK/STAT signaling pathway to avoid cross-talk between cells.
- Preventive: it decreases the mRNA synthesis of 12 major cytokines and chemokines to promote a prolonged soothing effect.

A clinical study on 25 volunteers has demonstrated a significant reduction of skin irritation and dehydration instantly in only 15 minutes.

### Instant decrease of skin inflammation

1. Cytokine binds to its receptor
2. Phosphorylation of receptor residues by JAK
3. Phosphorylation of STAT
4. Dimerization of STAT and entrance in the nucleus

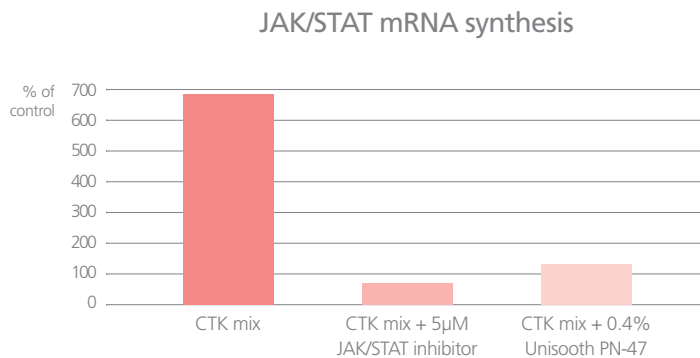


# Biological activity

## Curative action: control of JAK/STAT, the major inflammation pathway (*in vitro* test)

Unisoath PN-47 was tested for its ability to inhibit the JAK/STAT pathway. Normal human epidermal keratinocytes (NHEK) were pre-incubated for 6 hours in a medium containing Unisoath PN-47 at 0.4% or the JAK/STAT inhibitor Pyridone 6 at 5µM. They were then treated with a cytokine mix (oncostatin M, Interleukin 17 and TNF-α) referred to as CTK mix for 24 hours. mRNA expression was determined by q-RT-PCR.

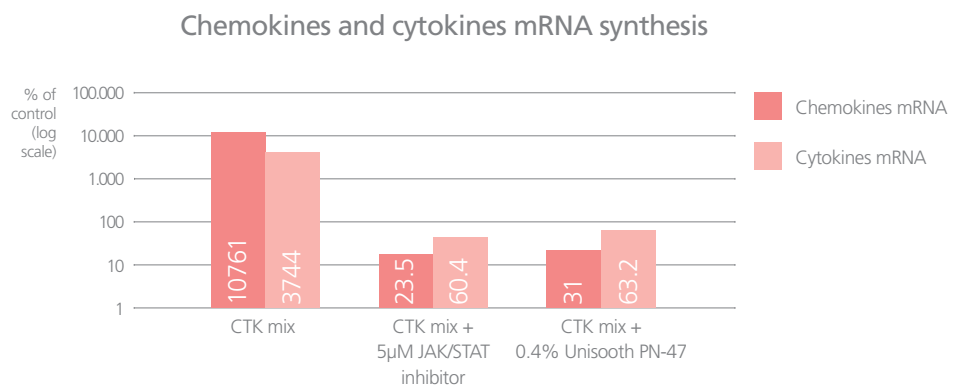
**Result:** Treatment with 0.4% of Unisoath PN-47 inhibits 80% of the JAK/STAT signaling compared to CTK mix to reach nearly the same level as Pyridone 6, a powerful reference JAK/STAT inhibitor.



## Preventive action: control of the expression of pro-inflammatory factors (*in vitro* test)

Unisoath PN-47 was tested for its ability to decrease chemokines (CCL5, CXCL10, CXCL5, CXCL6) and cytokines (IL-1α, IL-1 receptor type 1, IL-1 receptor type 2, IL-1 receptor antagonist, IL-4 receptor, IL-8 and TNF-α). Normal humal epidermal keratinocytes (NHEK) were pre-incubated for 6 hours in a medium containing Unisoath PN-47 at 0.4% or the JAK/STAT inhibitor Pyridone 6 at 5µM. They were then treated with a cytokine mix (oncostatin M, interleukin 17 and TNF-α) referred to as CTK mix for 24 hours. mRNA expression was determined by q-RT-PCR.

**Result:** Treatment with Unisoath PN-47 at 0.4% inhibits cytokines and chemokines mRNA synthesis by respectively -98% and -99% compared to CTK mix.



# Efficacy

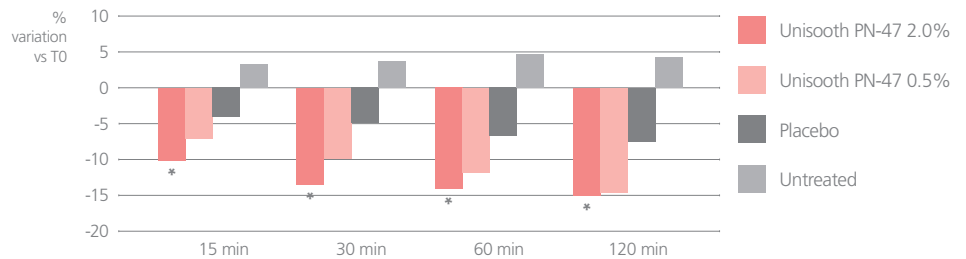
## An instant effect on skin irritation (Clinical efficacy)

Unisoath PN-47 was tested versus a placebo at 0.5% and 2% for its ability to instantly reduce skin irritation. Skin irritation was induced by a 24h patch occlusion with a 2% sodium lauryl sulfate (SLS) aqueous solution on the skin. Erythema index (Mexameter® MX 18) and Trans-Epidermal Water Loss (Tewameter® 300) were evaluated after 15 min, 30 min, 60 min and 120 min.

**Result:** Treatment with a cream containing 0.5% and 2% of Unisoath PN-47 reduces SLS-induced TEWL. After 15 min, Unisoath PN-47 at 2% decreases epidermis dehydration by -10%, and by -15% in 2 hours.

\*p<0.05 compared to placebo, Student's t Test

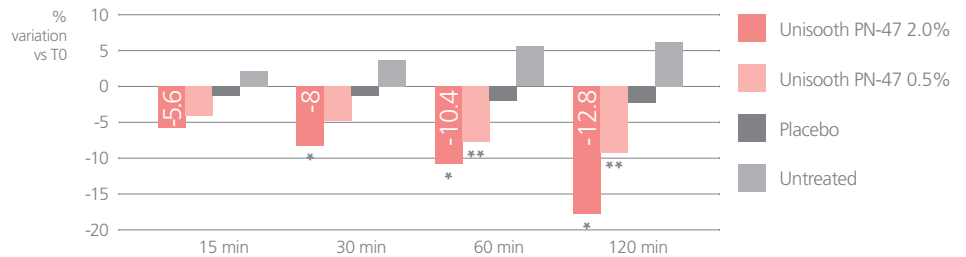
Reduction of epidermis dehydration (TEWL)



**Result:** Treatment with a cream containing 0.5% and 2% of Unisoath PN-47 reduces significantly skin redness. After only 30 min with Unisoath PN-47 at 2% skin redness is significantly decreased by -8%, and by -13% after 120 min.

\*p<0.05/\*\*p<0.01 compared to placebo, Student's t Test

Instant soothing effect (skin redness)



# Summary

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## Technical information

|               |   |
|---------------|---|
| INCI:         | Panthenyl Triacetate, Naringenin  |
| Origin:       | Plant extract + Organic synthesis   |
| Preservation: | Preservative-free   |
| Appearance:   | Clear, yellow liquid  |
| Solubility:   | Soluble in oils, alcohols, glycols  |
| Dosage:       | 0.5-2%  |
| Processing:   | Can be added at the end of the formulation process under stirring or homogenizing.<br>It can be heated for a short time with the oil phase of a formulation.<br>Formulate at pH between 4.0 and 7.0 and temperature below 40°C. |

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## Claims

|               |  |
|---------------|--|
| Claims:       | Instant soothing, control of skin irritation reactivity, daily protection against aggressions  |
| Applications: | Sensitive skin care, calming cream, after sun lotion, after shave products, post depilatories cream, post laser spray, lotion for irritated scalp, cream for insect bite |

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