



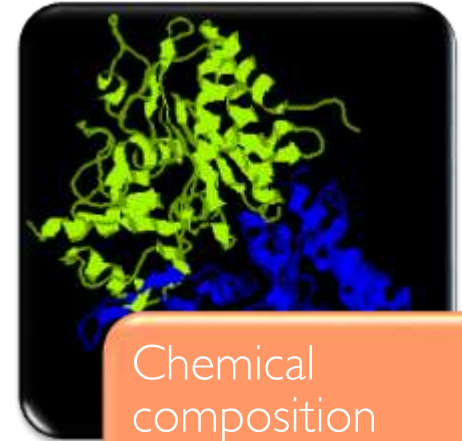
FruitPeel™

Glycerin based extract from Tropical Fruits standardized in carbohydrates and total alpha hydroxyacids (AHAs)

INCI name Suggested: Water (and) Glycerin (and) Spondias Mombim Pulp Extract (and) Mangifera Indica (Mango) Pulp Extract (and) Musa Sapientum (Banana) Pulp Extract

From where the idea came out?

FruitPeel™



Cell turnover and new antiageing

- Mild cell turnover stimulation
- Face and skin adipocytes filling

Tropical fruits

- Banana
- Caja
- Mango

Chemical composition

- AHAs from fruits
 - Citric, malic, glycolic
- Sugars, carbohydrates, minerals

Marketing Claims

- Mild cell turnover
- AHAs naturally found in tropical fruits
- Different mechanism of action from traditional AHAs
- Accelerates cell turnover without excessive discomfort
- Keratolytic effect, reinforcing the cutaneous barrier due young corneocytes cohesion
- Intelligent lipogenic effect, stimulating silent aged genes
- Enhance general features of cutaneous relief

Ageing: changes in cutaneous tissue

Epidermis

- Keratinocytes number decrease
- Turnover cell reduction
- Tapering
- Corneocytes size increase
- Intercellular lipids reduction
 - Ceramides, cholesterol, fatty acids
 - Susceptibility and transepidermal water loss increase (TEWL) → dehydration
- Melanocytes: number reduction; size and activity



Ageing: changes in cutaneous tissue

Dermis

- Fibroblast number and activity reduction
 - ↓ collagen, elastin and glycosaminoglicans
- Collagen loses solubility
- Elastin (elastic tissue) → degradation and elasticity loss
- Photolytic enzymes activity increase and inhibitors activity reduction
 - Metaloproteinases Matrix (MMP's)
 - Metaloproteinases (TIMP's) tissue inhibitors
- Dermal-Epidermal Junction (DEJ) decrease

Ageing: changes in cutaneous tissue

Micro vein

- Changes in blood vessels and veins
 - oxygen offer and essential nutrients reduction to dermis and epidermis
 - whiteness increase and temperature decrease

Sebaceous and sweat glands

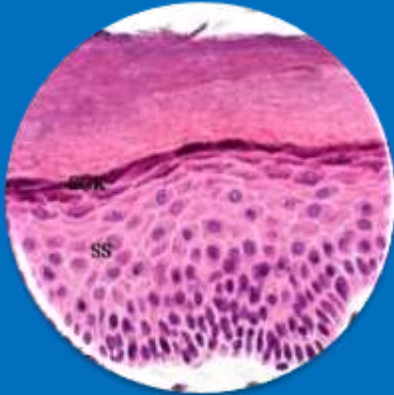
- Hormonal stimulation decrease → ↓ secrete activity
- Decrease in ecrin glands number and apocrin glands function

Ageing: changes in cutaneous tissue

Hypodermis (subcutaneous tissue)

- Adipocytes waste away
 - face, legs, hands and feet
 - contribution to wrinkles formation
- Proteins genic activity and adipocytes transcription factors reduction
 - ↓ SREBP-1
 - ↓ intercellular lipids buildup





Epidermis

- Transglutaminase-1
- Claudin-1 (Tight Junction)



Dermis

- Procollagen



Hypodermis

- SREBP-1
- Intracellular triglycerides

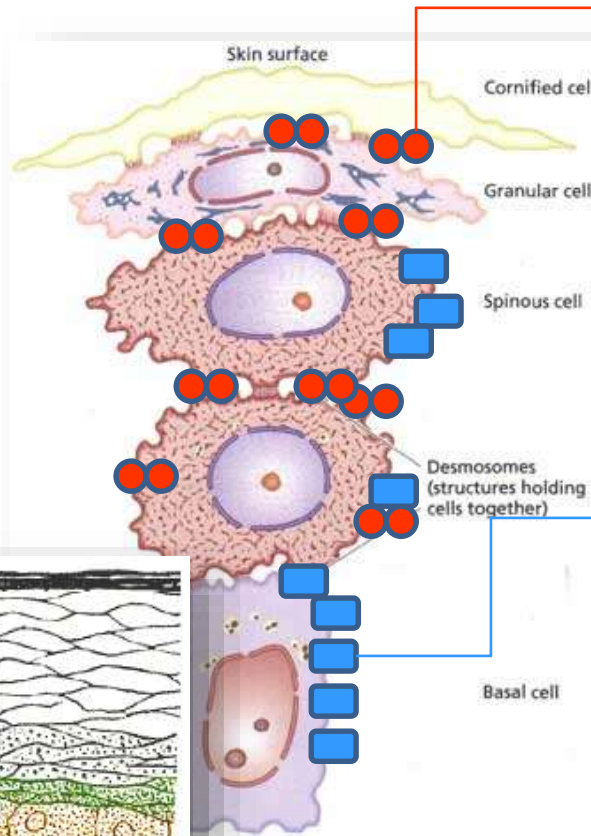
FruitPeel™

Mechanisms of Action definition



Epidermis

- TGase-I
- Claudin-I

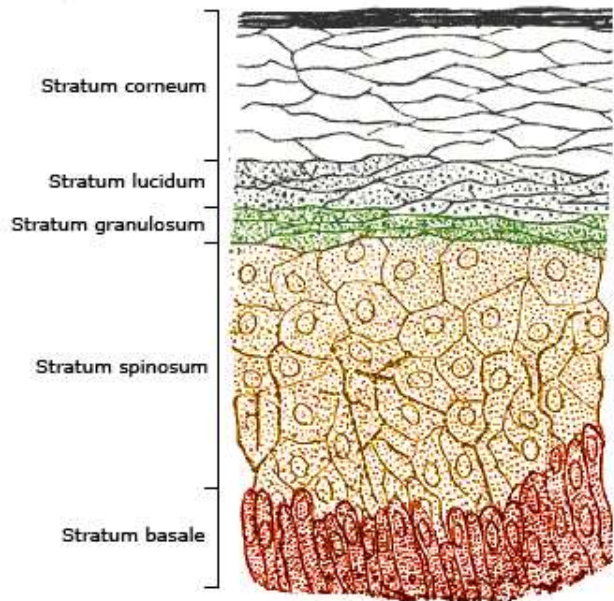


Transglutaminase (TGases)

Calcium dependent enzyme that catalyzes covalent bond among proteins and peptides from SC

Claudin-I

Structural tight junction (TJ) protein essential to adherence keratinocytes. Indicates cell turnover index



Traditional AHAs

Chemical aggression exfoliation ($\text{pH} \leq 3,8$)

Chemical aggression = cell proliferation (turnover)

As consequence...



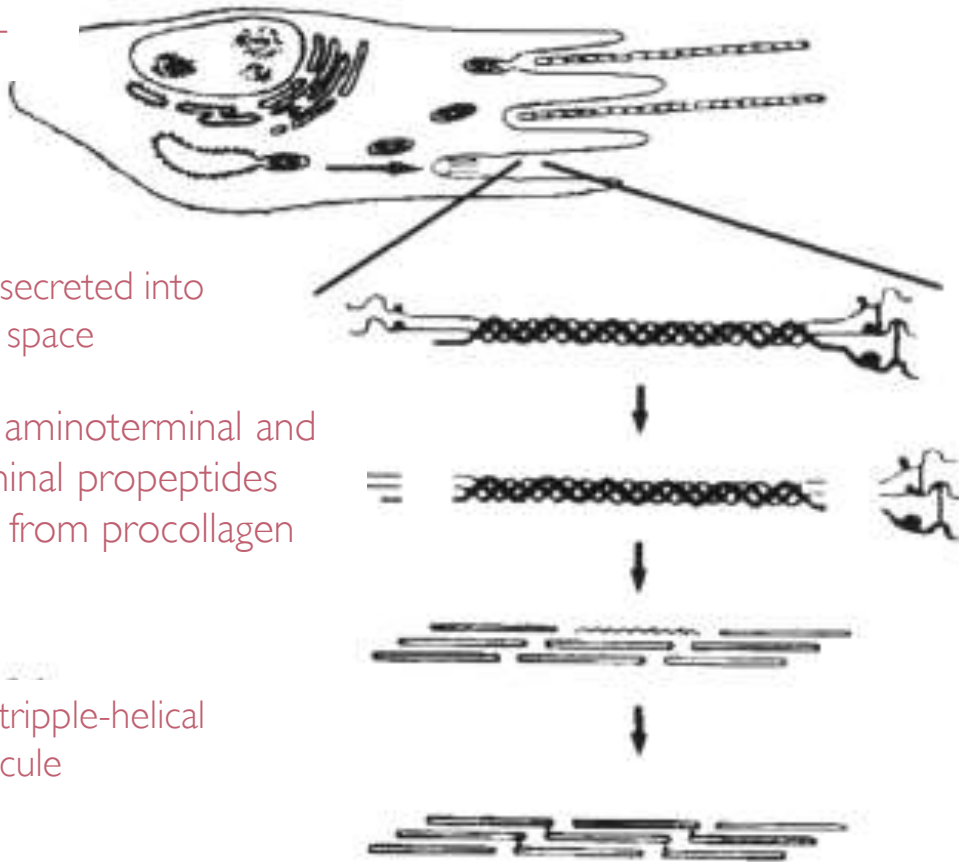
FIBROBLAST



Procollagen secreted into extracellular space

Cleavage of aminoterminal and carboxyterminal propeptides
propeptides from procollagen molecule

Formation of tripple-helical collagen molecule



Dermis

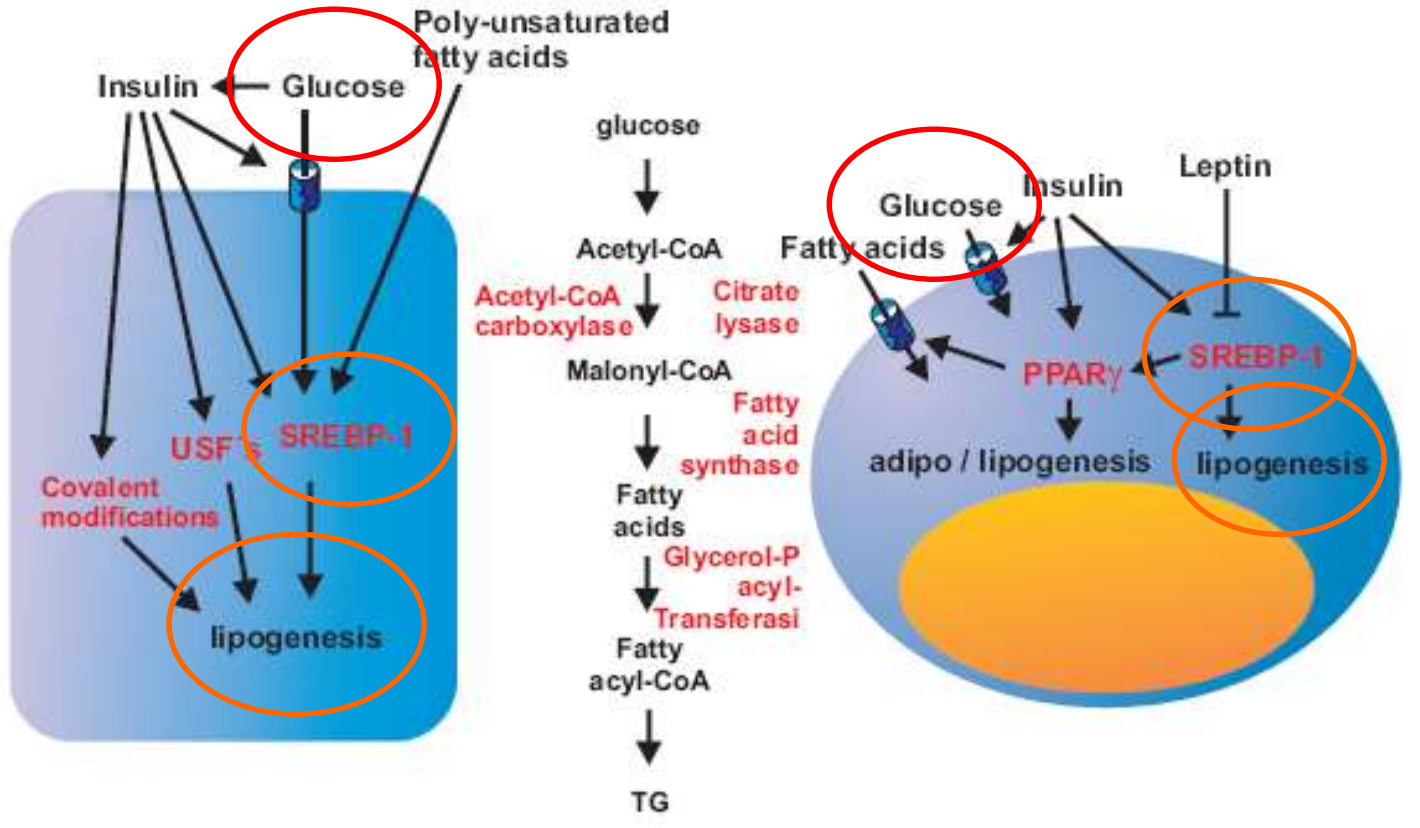
- Procollagen

Procollagen = collagen again, functional collagen



Hypodermis

- SREBP-1
- Triglycerides



Summary:

- Codifying gene silencing for SREBP-1 due to ageing
- \downarrow SREBP-1 = \downarrow Lipogenesis = \downarrow intercellular TAGs = \downarrow tissue filling

Safety and Efficacy Assessment
FruitPeel™

Safety

In vitro

Clinical

Citotoxicity

Phototoxic Potential

Iritant Potential

Iritant Potential

Allergenic Potential

Efficacy

Clinical

In vitro

Epidermal action

Dermal action

Hypodermal action

Immunohistochemical expression and localization of Claudin-1

TGase I activity

Procollagen gene expression

SREBP-1 gene expression

Lipolysis index reduction

Safety

In vitro

Cytotoxic Potential (XTT Test)

Citotoxicity not relevant

Phototoxicity (3T3-NRU)

Non-phototoxic

Irritant Potential (HET-CAM Test)

Weakly irritant

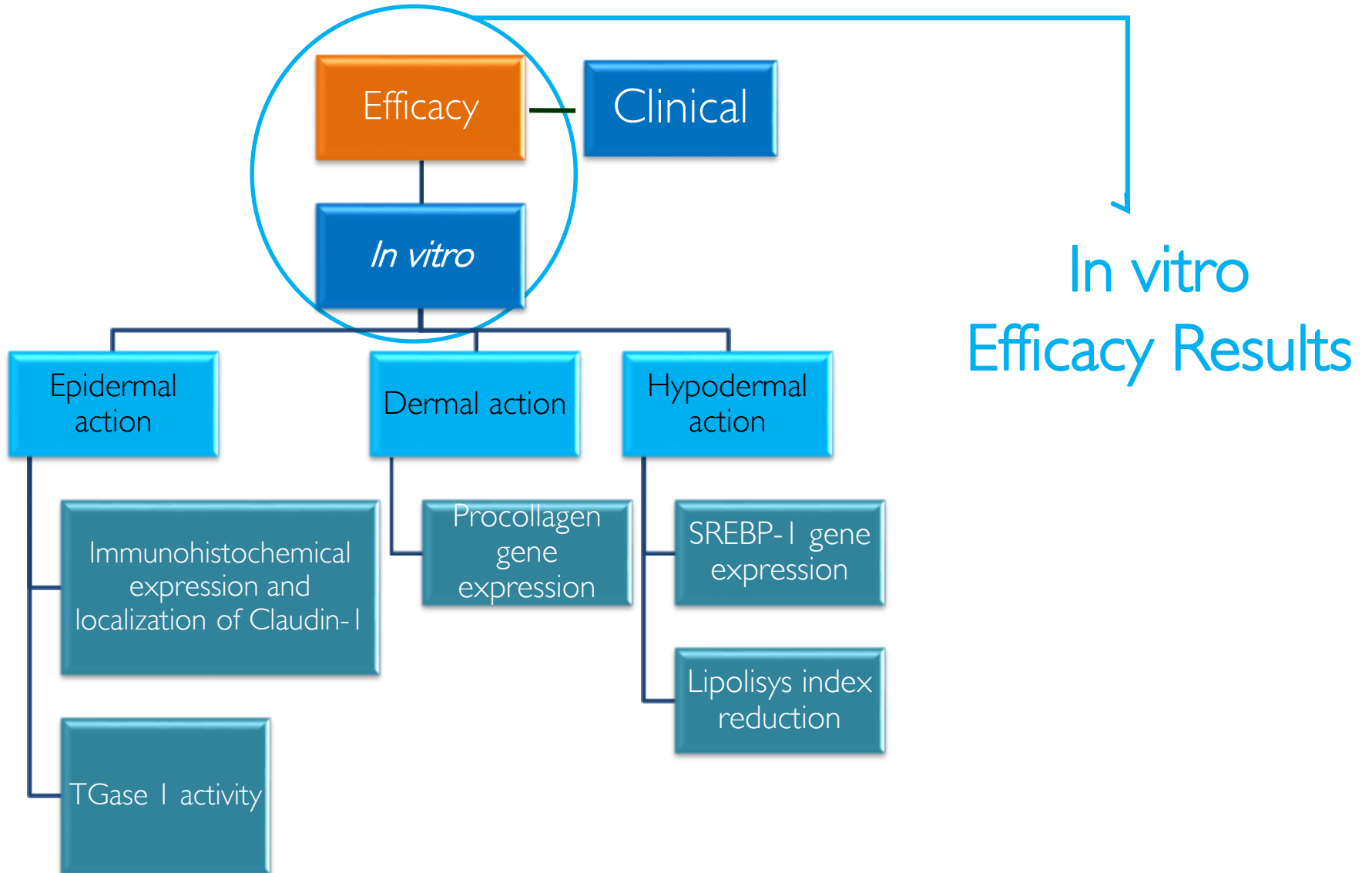
Clinical

Primary skin irritation and cumulative (Open Test, PC5)

Good skin compatibility

Allergenic potential (HRIPT)

No allergenic potential

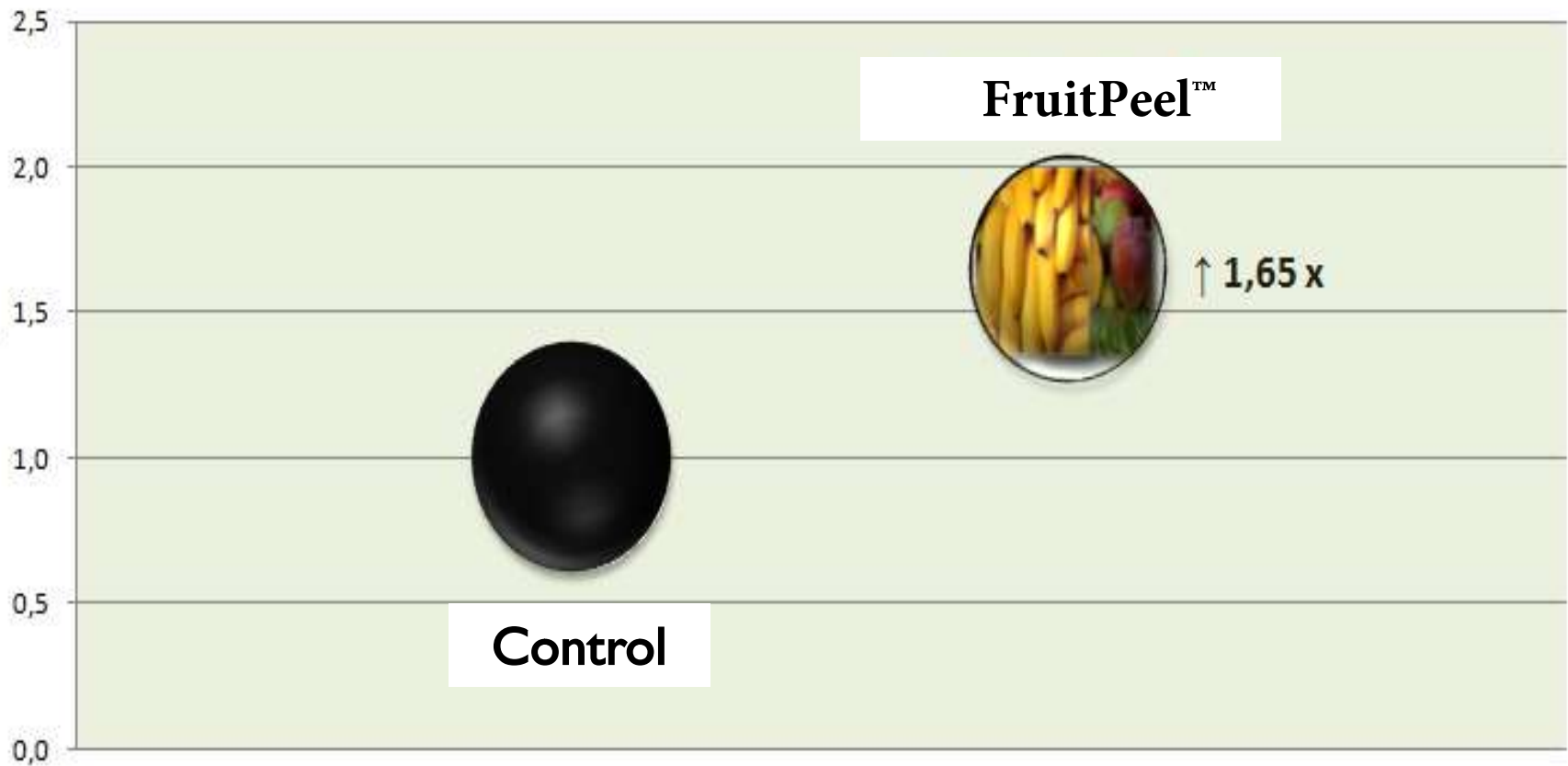




Epidermal Action

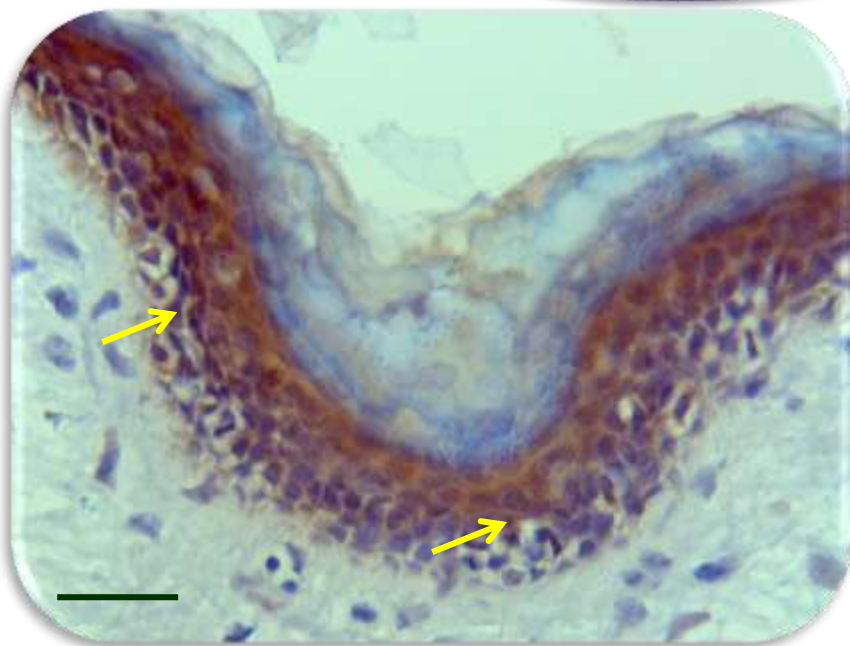
Claudin-1
Expression

CLAUDIN 1 (mRNA) gene expression,
related to control

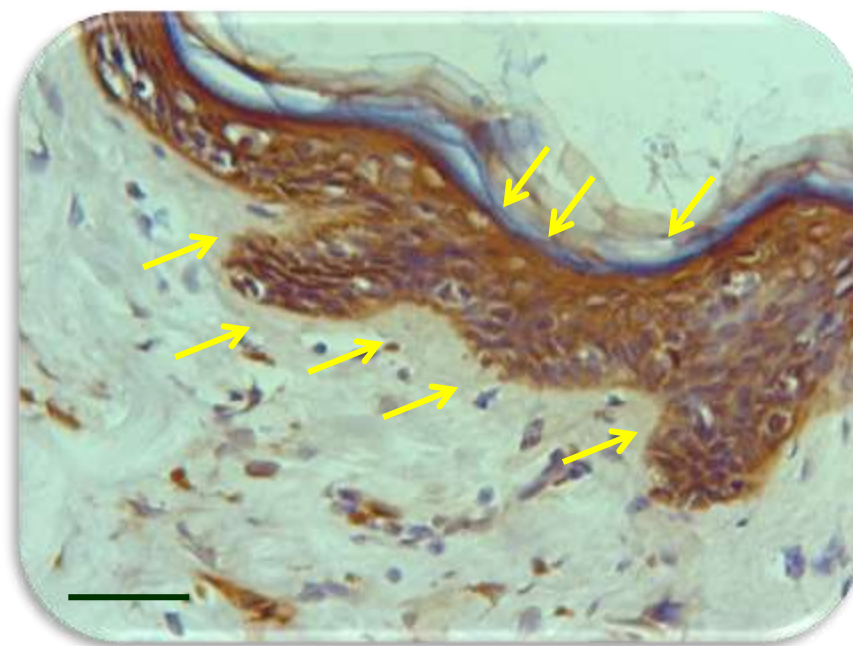


Epidermal Action

Claudin-I
Immunohistochemical



Control (40x)



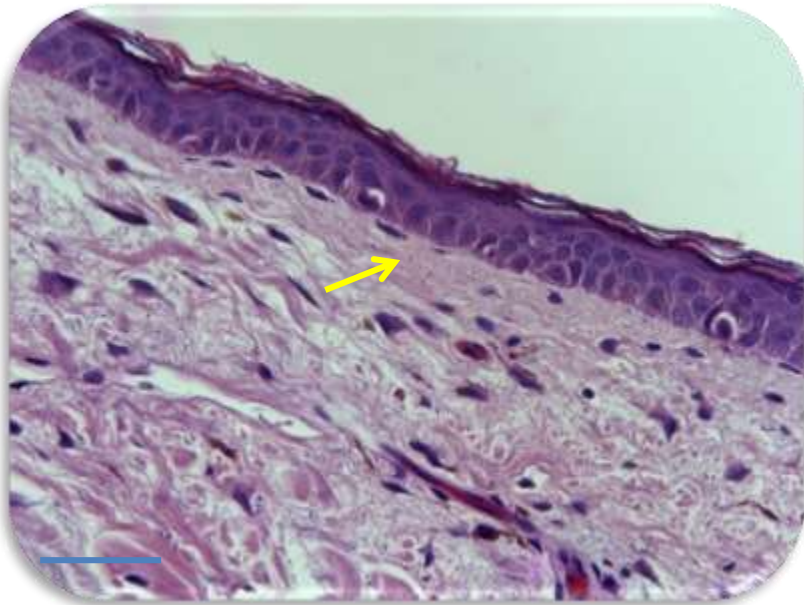
FruitPeel™ 0,04% (40x)

25µm

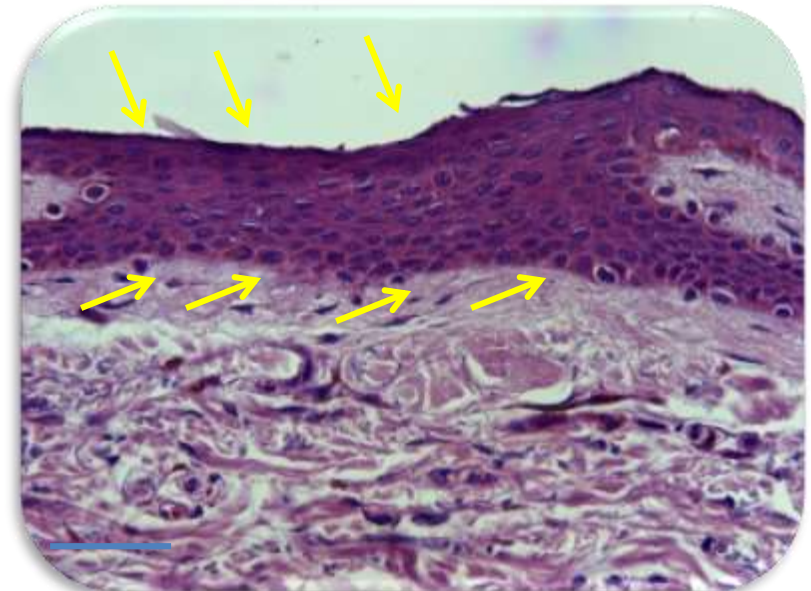
Intensity and distribution for brown area indicate Claudin-I expression

Epidermal Action

Skin general condition (HE)



Control



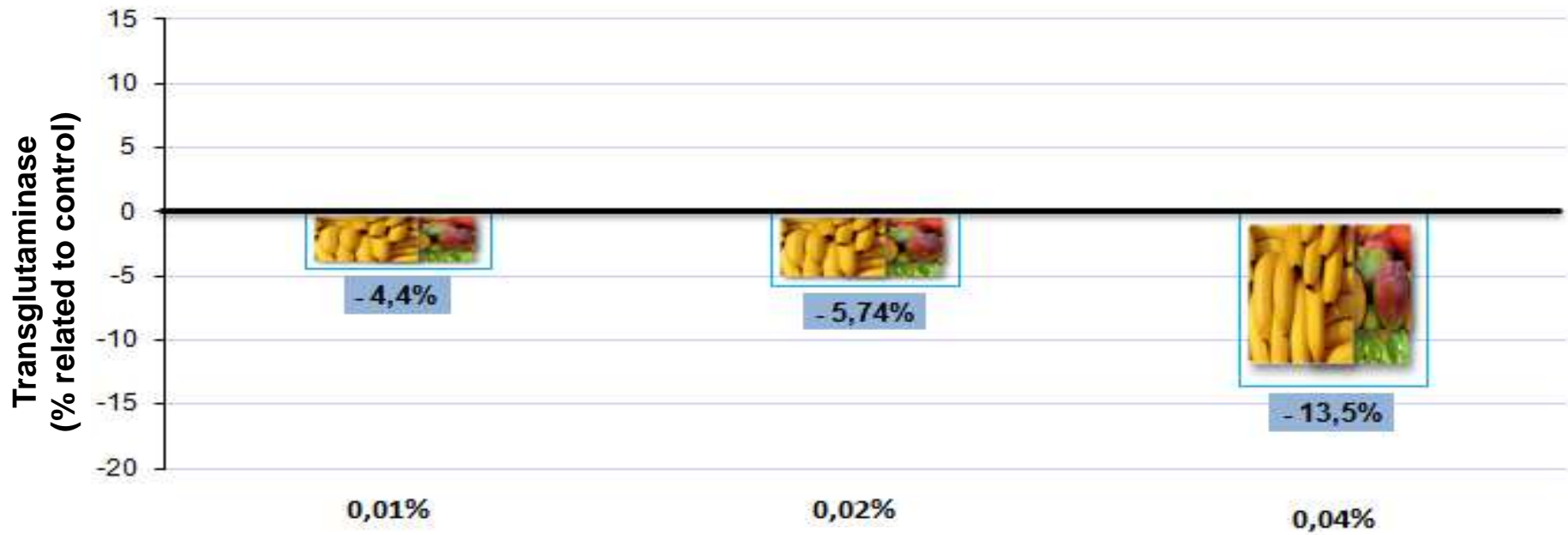
FruitPeel™ 0,04%

25µm

Epidermal Action

TGase I

ELISA/colorimetry



FruitPeel™

Dermal
Action

Procollagen
expression

PCR-Real Time

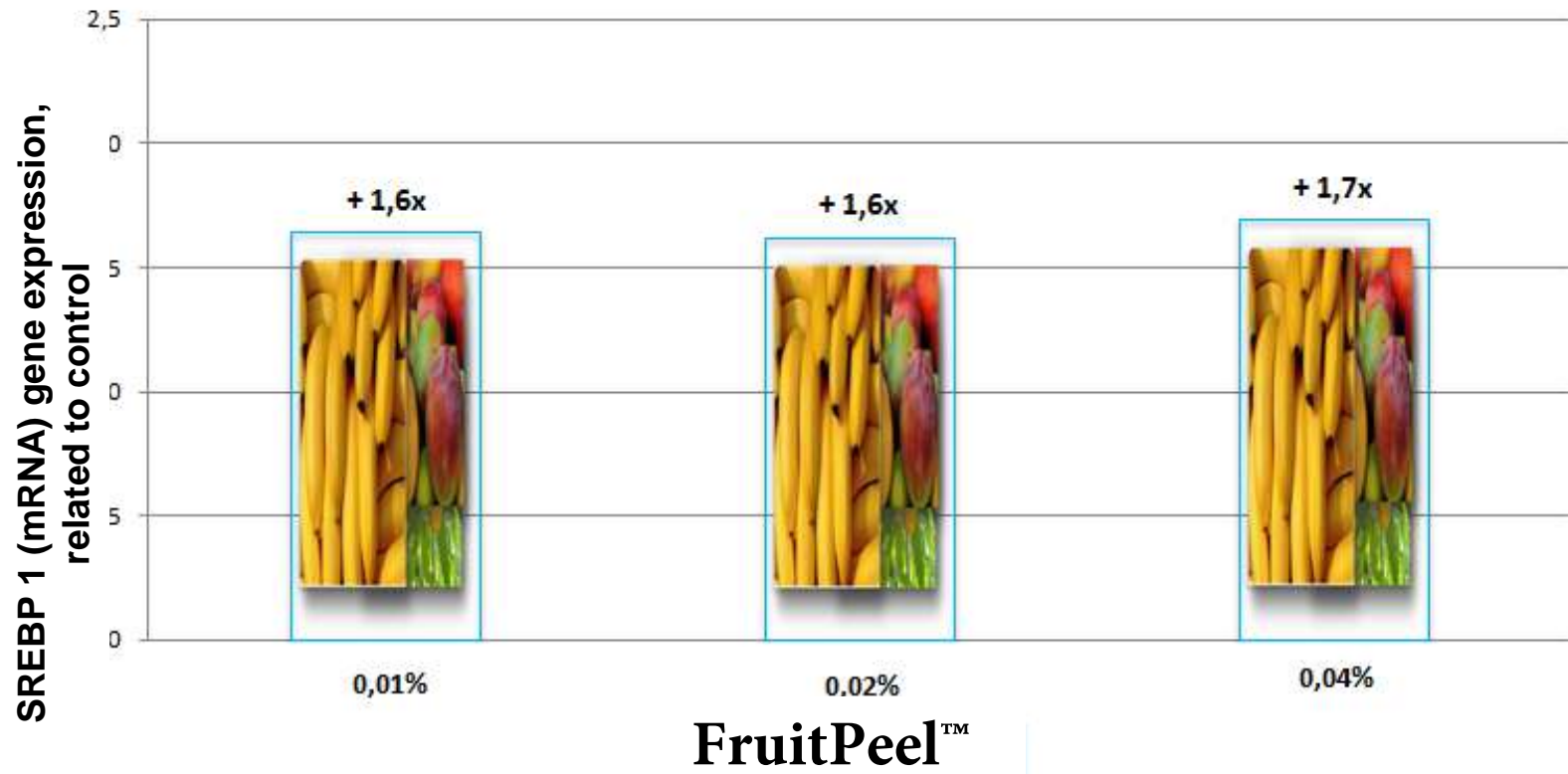


FruitPeel™

Hypodermal
Action

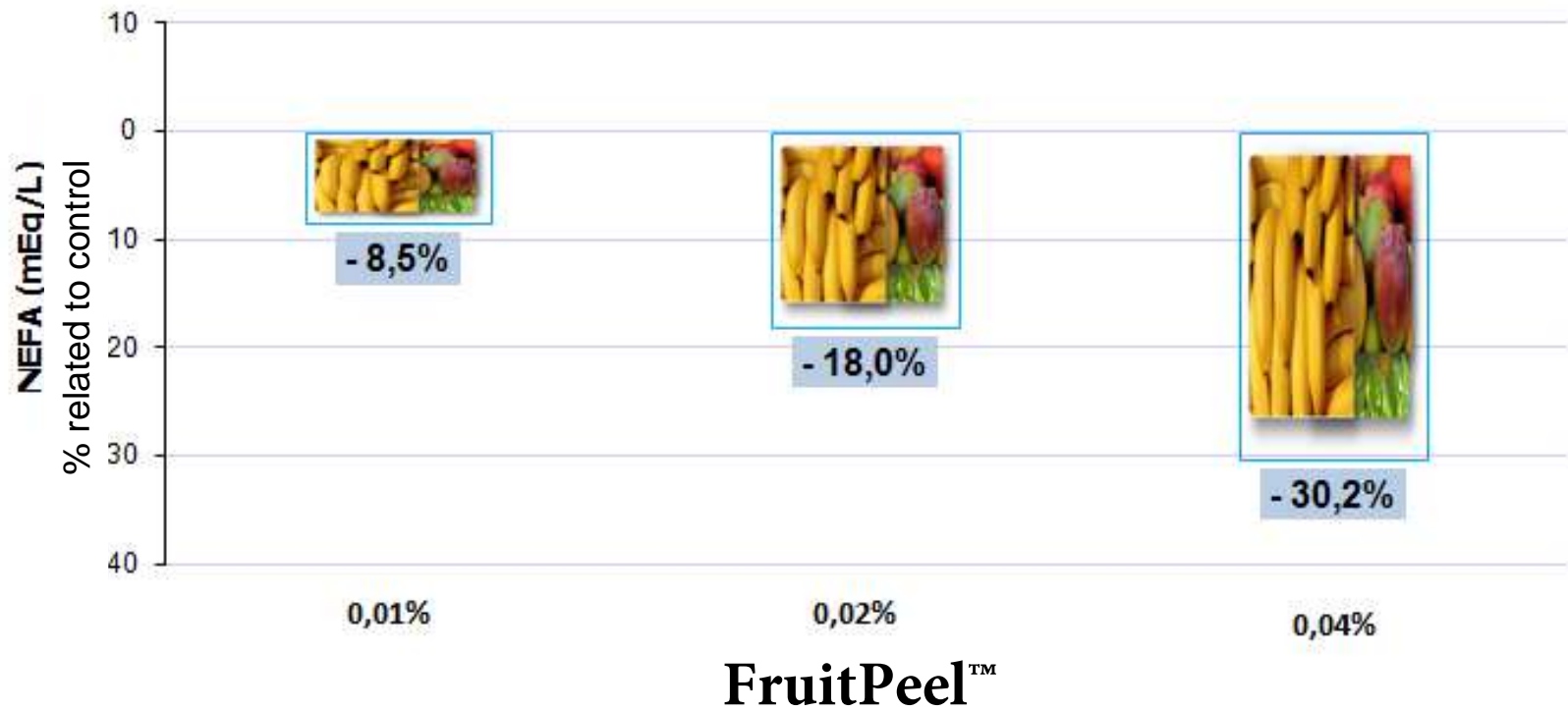
SREBP-1
expression

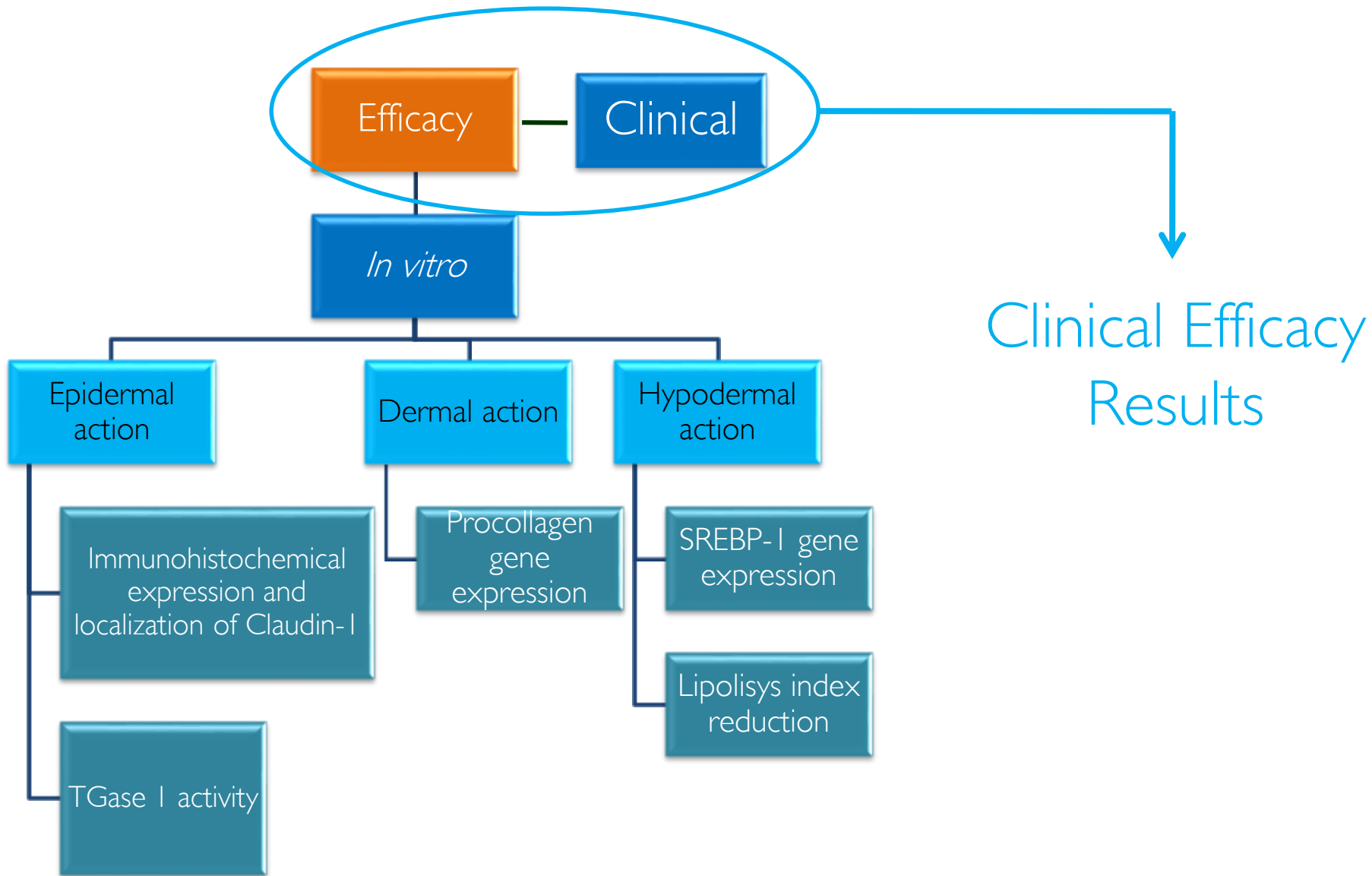
Aged adipocytes due to hormonal deficit



Hipodermal
Action

Intracellular
TAGs buildup





Clinical Assessment

Subjective Efficacy –

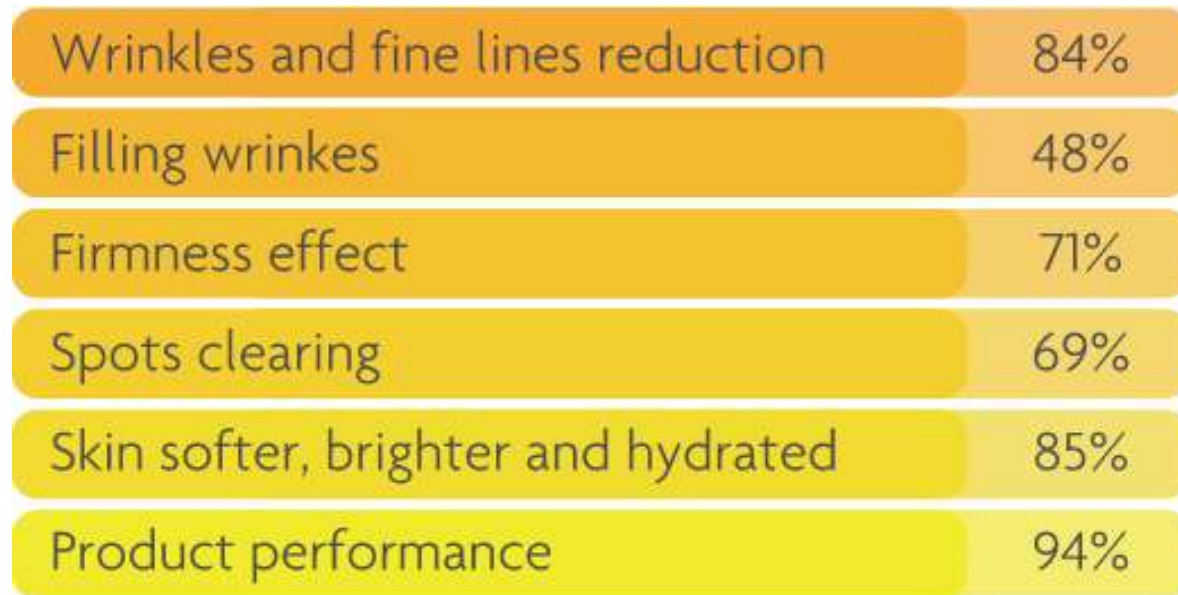
Sensory evaluation and Cosmetic Appreciability

- 35 volunteers – aged between 35 and 55 years old
- 28 days treatment, under normal use conditions
- Placebo cream and cream with **FruitPeel™** 3%



Clinical Assessment

Subjective Efficacy



Why to use FruitPeel™?

- **FruitPeel™** is a non aggressive cell turnover
- **FruitPeel™** contains natural AHAs from tropical fruits
- **FruitPeel™** accelerates epidermal cell turnover by promoting balance among main cell structures adherence

Why to use FruitPeel™?

- **FruitPeel™** promotes keratolytic effect, reinforcing the barriers due to more young corneocytes cohesion
- **FruitPeel™** has stimulator effects in wrinkles and tissues filling, due to its capacity to regulates gene activity related to adipogenesis
- **FruitPeel™** promotes triple benefits to the skin, acting on epidermis, dermis and hypodermis