



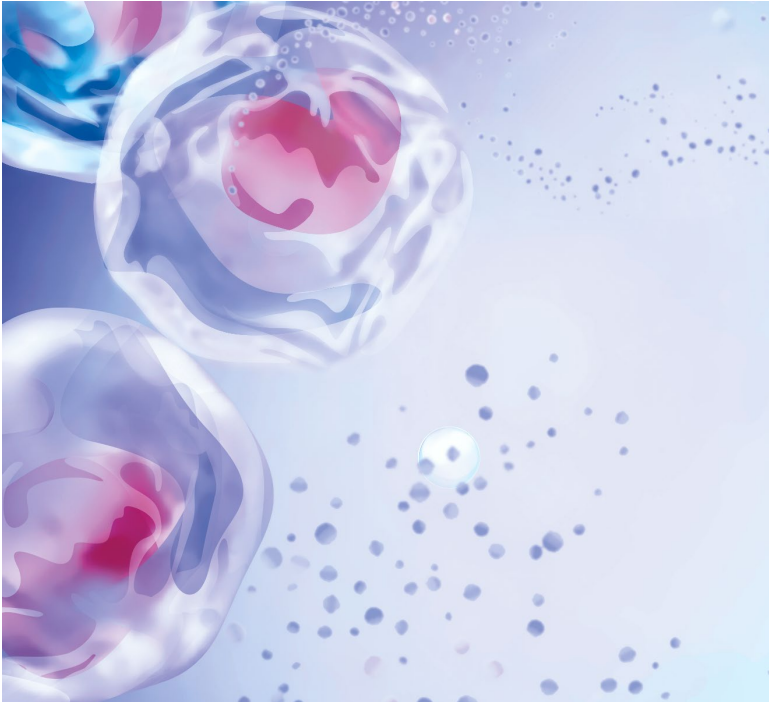
PhytoCellTec®

PhytoCellTec™ Exosomes

Dual exosome power for rejuvenation



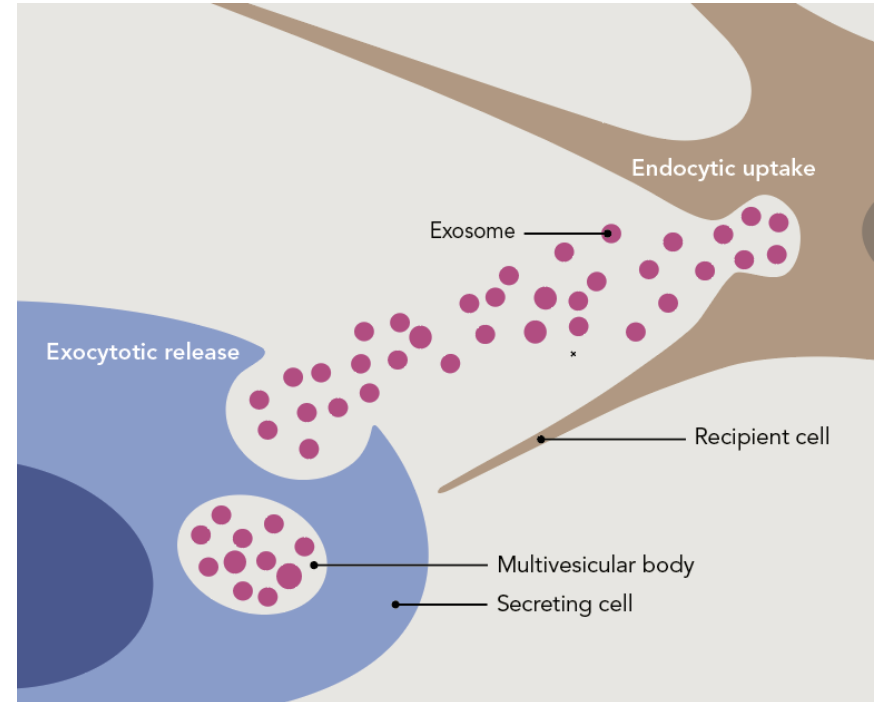
Exosomes – a Trending Topic with Real Potential



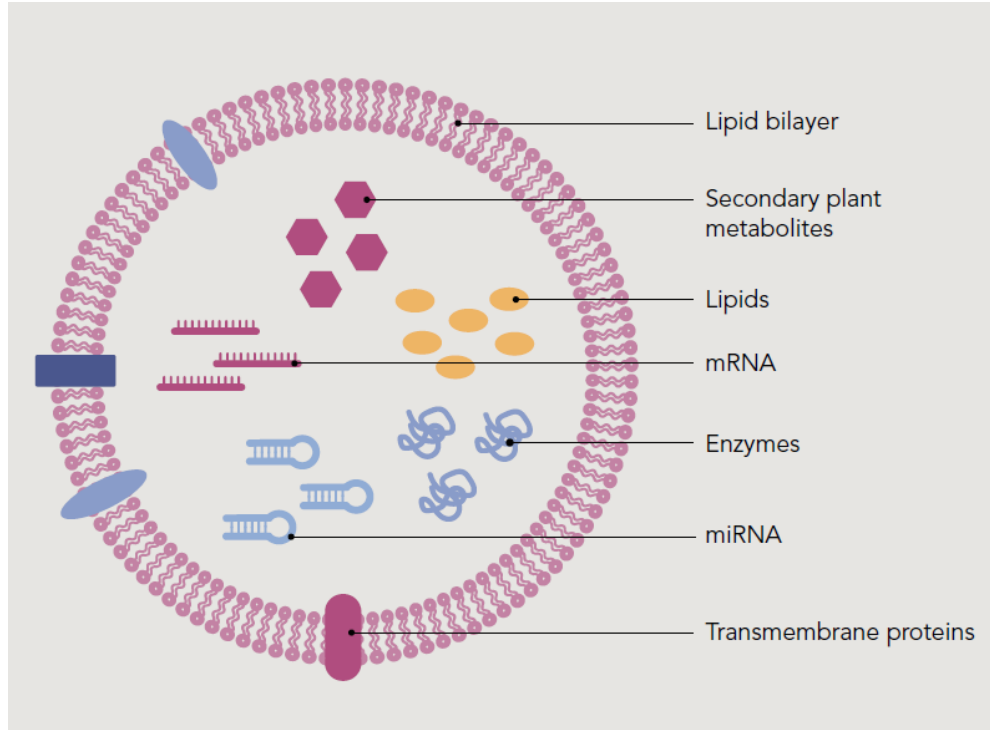
- Exosomes have been increasingly discussed in the past few years due to their potential for the **diagnosis** and therapy of **diseases**, such as cancer and neurodegenerative diseases.
- Cutaneous **medical aesthetics** use effect of exosomes on wound healing, skin pigmentation, and hair loss.
- In the meantime, exosomes have also evolved into a hot topic in the world of **cosmetics**.

What is the Function of Exosomes?

- Exosomes are an important component of cellular **communication** systems.
- Exosomes released by one cell are e.g. **taken up by the recipient cell** (endocytic uptake) → response in this cell is initiated.
- Signaling via exosomes plays a **major role** in many **cellular processes**, such as signal transduction and immune response.



Plant-Derived Exosomes



- Similar to human cells, plant cells use extracellular vesicles to transfer messages from cell to cell (RNAs, proteins, lipids, and secondary metabolites).
- Plant-derived exosome-like nanovesicles → **“plant exosomes”**
- Function in the plant: intercellular communication, defense system or cell wall organization.
- **Similar composition, size and function** as in skin tissue.

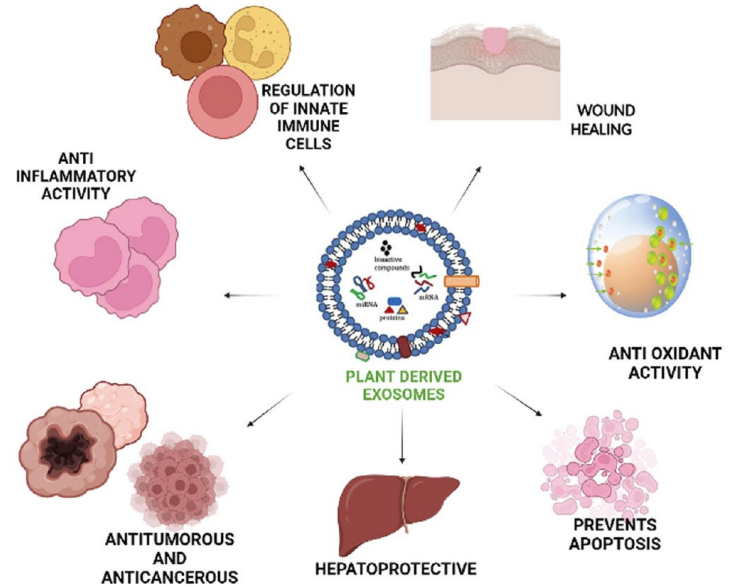
Plant-Derived Exosomes

- Plant-derived exosomes can be **taken up by mammalian cells** and have a beneficial effect on human health!
- Exosomes from different plants showed anti-inflammatory, antioxidant, anti-tumor, or wound healing potential in human (1, 2).
- Recent research results confirmed the **anti-aging** potential of **plant exosomes** for the skin (3).
- **Plant exosomes** can be taken up by the **skin** where they unleash their full potential.

(1) Subha, D. et al. (2023) Plant Nano Biology 3:100022.

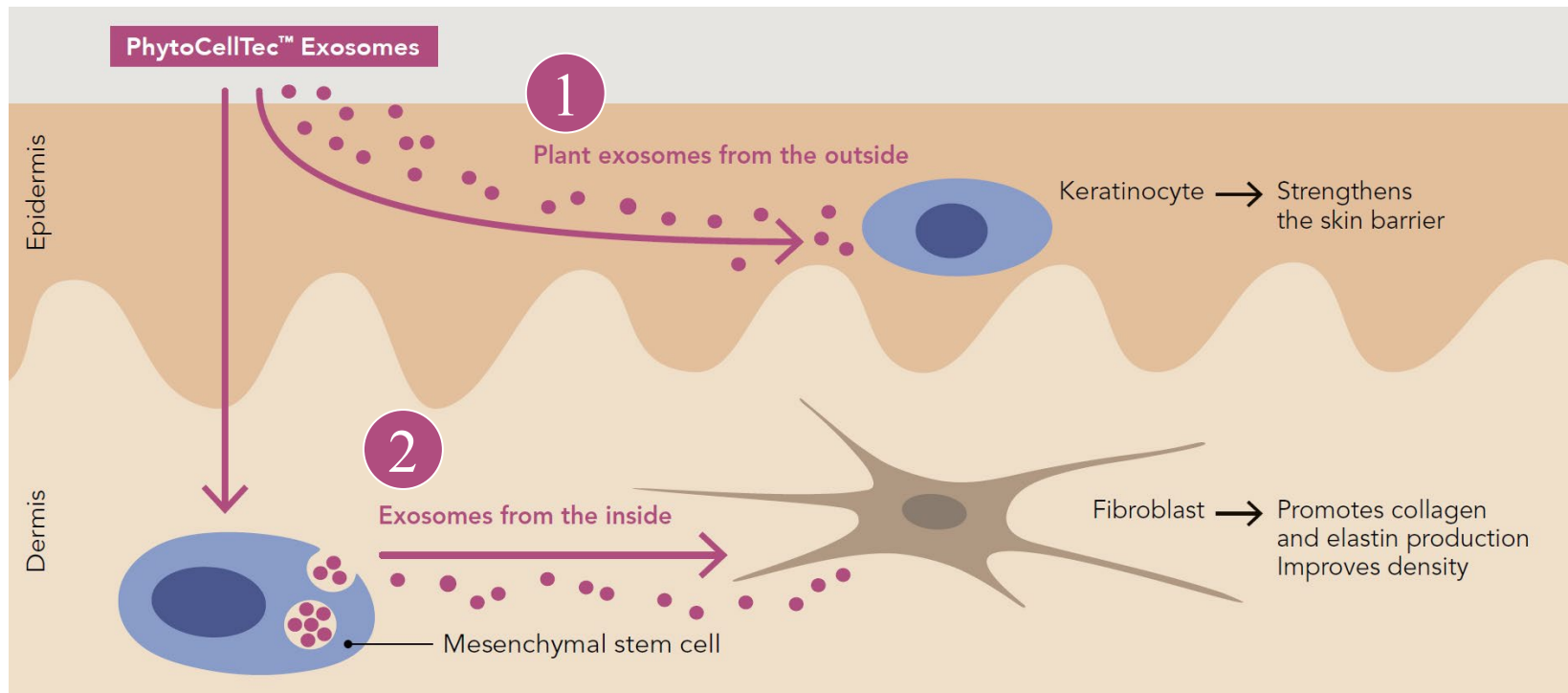
(2) Mu, N. et al. (2023) Int J Nanomedicine 18:4987 – 5009.

(3) Trentini, M. et al. (2022) Cells 11:3950.



*Schematic representation of different therapeutic potential of plant exosomes
D. Subha, K. Harshni, K.G. Madhikiruba et al. Plant Nano Biology 3 (2023) 100022*

PhytoCellTec™ Exosomes: A Dual Activity to Rejuvenate the Skin



Dual Approach of PhytoCellTec™ Exosomes

1. Plant exosomes from the outside

- Goji stem cell-derived exosomes act directly on the outer cell layer, the **keratinocytes**, to strengthen the barrier and to protect skin from aggressive external factors and dehydration.

→ **Strengthened skin barrier, reduction of fine wrinkles.**

2. Endogenous exosomes produced by mesenchymal stem cells

- Increased production of exosomes mesenchymal stem cells (MSCs) release
- Fibroblast in the dermis are activated through exosomes to produce more collagen and elastin

→ **Production of extracellular matrix, increase of skin density, reduction of sagging skin, V-shaped face and lifted breasts.**

Stem Cells from Goji Plant Contains Valuable Exosomes



- Bush up to 3 m in height native to Southeast Europe and Asia.
- Belongs to the nightshade family such as potatoes and tomatoes.
- Legend: Goji berries were first discovered by a Buddhist monk. The monks who incorporated them into their diets lived longer than those who didn't.
- Goji berries are considered superfruits / health food.
- Food supplements claiming anti-aging, immune boosting, energizing improvement of memory.
- Recent studies demonstrated that exosomes from goji plants provide health benefits (muscle atrophy and spinal cord injuries)*.

*Zhou, X. et al. (2024) J Nanobiotechnology 22(1):276. Wang, Q. et al. (2024) Bioeng Transl Med. 9(4):e10646.

Plant Stem Cells are an Ideal Source for Exosomes



It is **difficult** to extract pure exosomes from plants due to the cell walls and intricate structure of plant cells!

- **Plant stem cells** are an ideal **source** for exosomes
- No **rigid cell walls**, no chlorophyll, have a high number of exosomes inside and around the cells.
- Additionally, they provide the **positive effects** of **plant stem cells** on **skin stem cells**.

Plant Stem Cells for Skin Stem Cells

The stem cells of a goji seedling, obtained and cultivated by Mibelle Biochemistry's PhytoCellTec™ technology, are the basis of the active ingredient PhytoCellTec™ Exosomes.



Goji seedling



Stem cells on agar plate



Stem cell culture



Goji plant stem cell extract containing:

- **exosomes** from plant stem cells
- **epigenetic factors** that have a vitalizing effect on skin stem cells

PhytoCellTec™ Exosomes Composition

Lycium Barbarum Cell Culture Extract (dry)	0.8 %
Isomalt	93 %
Phospholipids	0.8 %
Aqua (residual moist)	~6 %

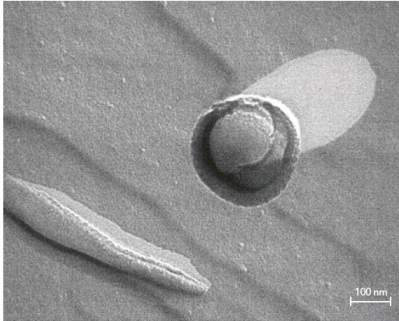
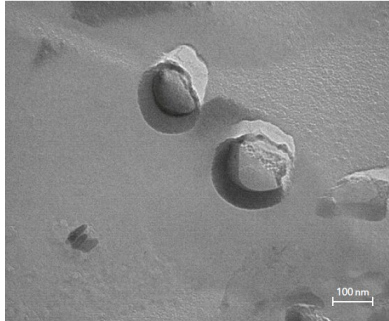
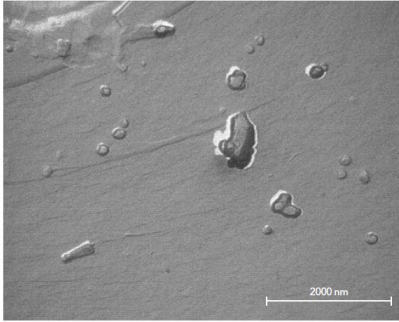
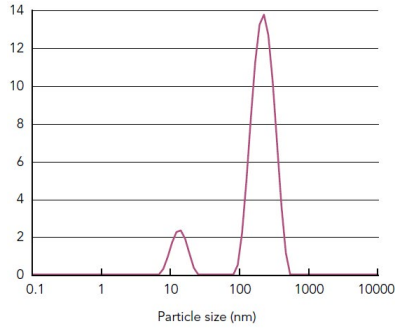
INCI (EU/PCPC) Declaration

Lycium Barbarum Callus Culture Extract (and) Isomalt (and) Lecithin (and) Aqua/Water

Recommended Use Level: 0.4 - 1 %



PhytoCellTec™ Exosomes Contains Exosomes



S-1301 / S-1305

Test sample:

Goji stem cell extract

Parameter:

- Vesicle size of exosomes
(Zeta Sizer Nano ZS from Malvern Panalytical)
- Pictures (freeze-fracture transmission electron microscopy)

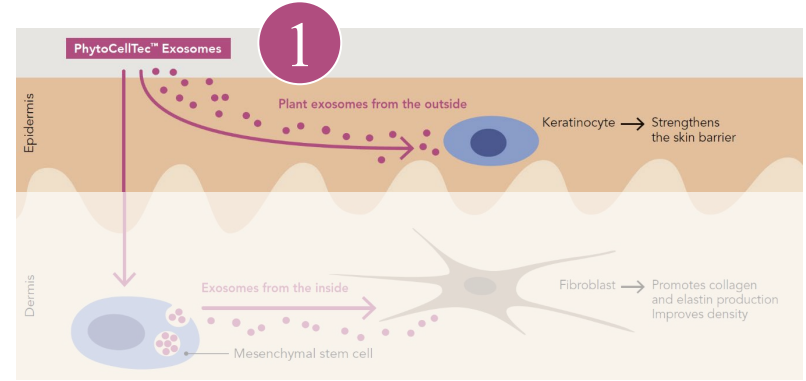


- PhytoCellTec™ Exosomes naturally contains **exosomes** derived from the **plant stem cells** of a **goji seedling**.
- Exosomes are of a size of approximately **200 nm**.
- **Images** confirmed the presence and the size of these exosomes that are enclosed by a **double-layer membrane**.



Plant Exosomes for Improved Skin Barrier Function

To assess the **first part of the dual action**, the effect of plant exosomes on the skin, gene expression in keratinocytes was analyzed.

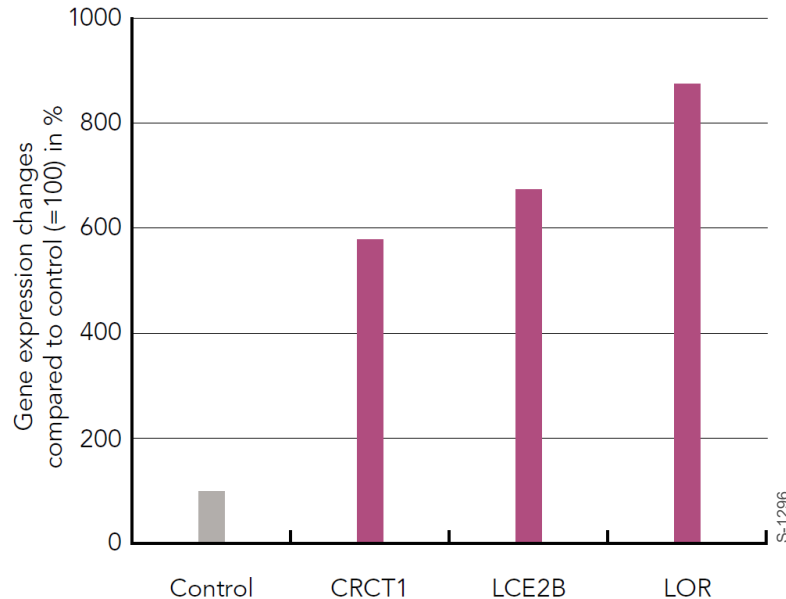


Cell type: Normal human epidermal keratinocytes (NHEK)
Test substance: 0.1 % Goji stem cell extract
Treatment: Application of 0.1% Goji stem cell extract to normal human epidermal keratinocytes for 24 hours

Parameter: Expression of genes important for the skin barrier and skin function (RT-qPCR)

Improved Skin Barrier Function

■ Control ■ 0.1 % Goji stem cell extract



Genes involved in the late **differentiation** and **cornification** process of keratinocytes were strongly upregulated (graph):

- CRCT1 = cysteine-rich C-terminal 1
- LCE2B = late cornified envelope 2B
- LOR = loricrin

Additionally, the gene expression of genes important for **regeneration** and **protection** was upregulated:

- epidermal growth factor (EGF, 108 %)
- fibroblast growth factor (FGF7, 161 %)
- heat shock protein (HSPA6, 407 %)
- antioxidant gene (SOD3, 64 %)

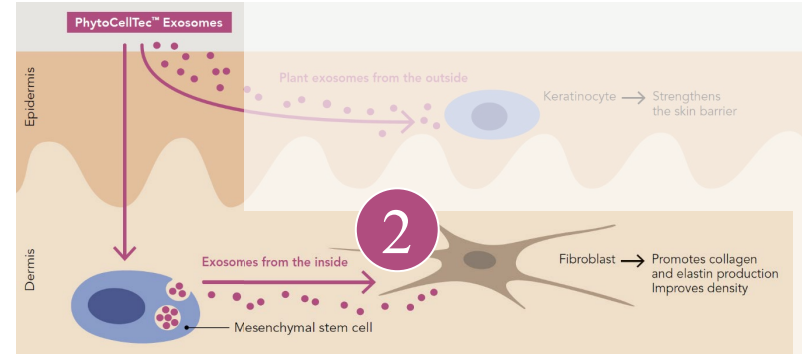


PhytoCellTec™ Exosomes supports the formation of the stratum corneum. Thereby, it can improve the function of the skin barrier to protect the skin from water loss and external aggressors.



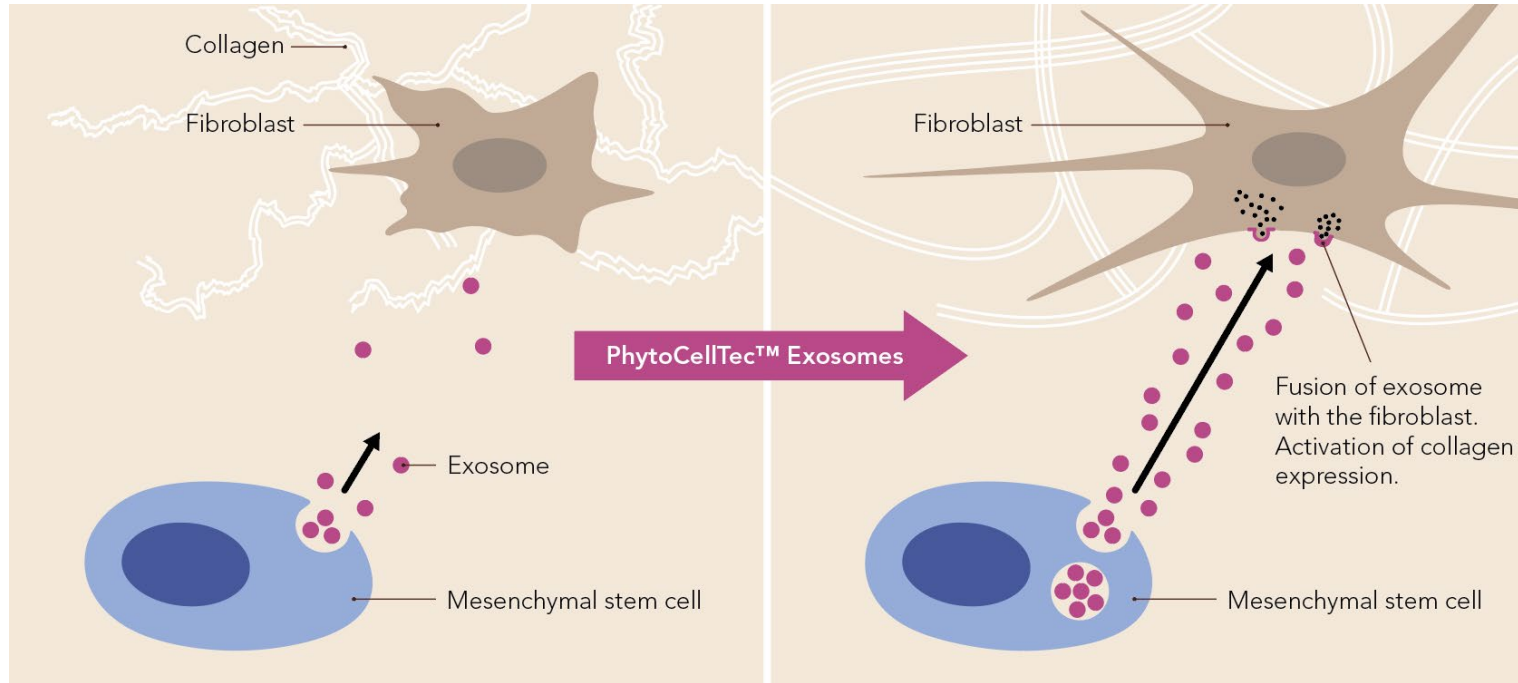
Activation of Aged Mesenchymal Stem Cells

To investigate the **second part of the dual action**, first, the effect of PhytoCellTec™ Exosomes on the vitality of mesenchymal stem cells (MSCs) was assessed.



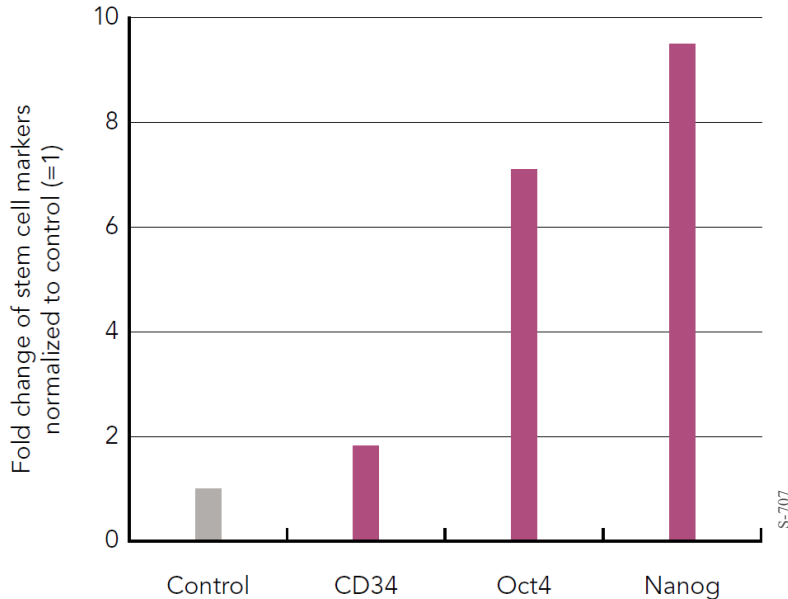
Cell type:	Adipose-derived human MSCs, grown for 14 passages to mimic the aging process.
Test substance:	1 % Goji stem cell extract
Treatment:	Aged MSCs +/- 1 % Goji stem cell extract Incubation for 72 h Control without treatment
Parameter:	Expression of stem cell markers (RT-qPCR).

Exosomes from Mesenchymal Stem Cells Rejuvenate Fibroblasts



Increase in Stem Cell Marker Expression

■ Control ■ 1 % Goji stem cell extract



Increased expression of different stem cell markers in aged MSCs.

→ PhytoCellTec™ Exosomes is able to **rejuvenate** mesenchymal stem cells and help them maintain their **stemness**.



Increase in Exosome Production by Mesenchymal Stem Cells

Assessment of the effect of PhytoCellTec™ Exosomes on the exosome production capability.

Cell type: Human mesenchymal stem cells (MSCs)

Test substance: 0.1 % Goji stem cell extract

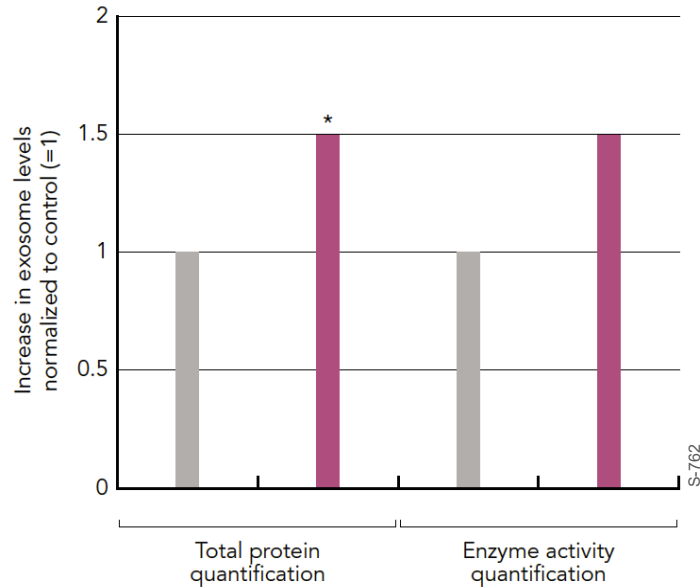
Treatment: Incubation of MSCs +/- (control) 0.1 % Goji stem cell extract for 24 h.
Isolation of exosomes that were released from the cells (EXO-prep kit)

Parameters: Quantification of exosomes by 2 methods:

- total protein amount (BCA protein assay)
- activity of acetylcholinesterase, a known exosomal protein (FluoroCet quantitation kit)

Increase in Exosome Production by Mesenchymal Stem Cells

■ Control ■ 0.1% Goji stem cell extract



*p<0.05 versus control



Both quantification methods revealed that treatment with Goji stem cell extract leads to an **increase in exosome production** by MSCs.



Stimulation of Extracellular Matrix Genes through Cell-Cell Communication

Investigation whether MSCs treated with Goji stem cell extract are able to **communicate** with **fibroblasts** to stimulate production of extracellular matrix (ECM) proteins.

Cell types:

Test substance:

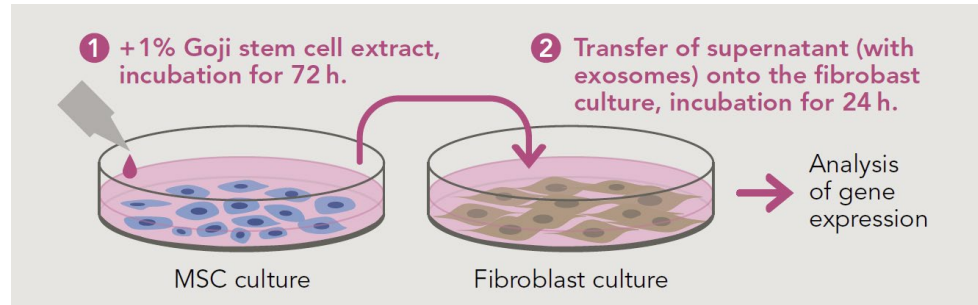
Treatment:

Human mesenchymal stem cells (MSCs), fibroblasts

1 % Goji stem cell extract

1. Treatment of MSC for 72 h with test substance

2. Add the supernatant (= conditioned medium) to the fibroblasts for 24 h (control: medium from untreated MSCs).

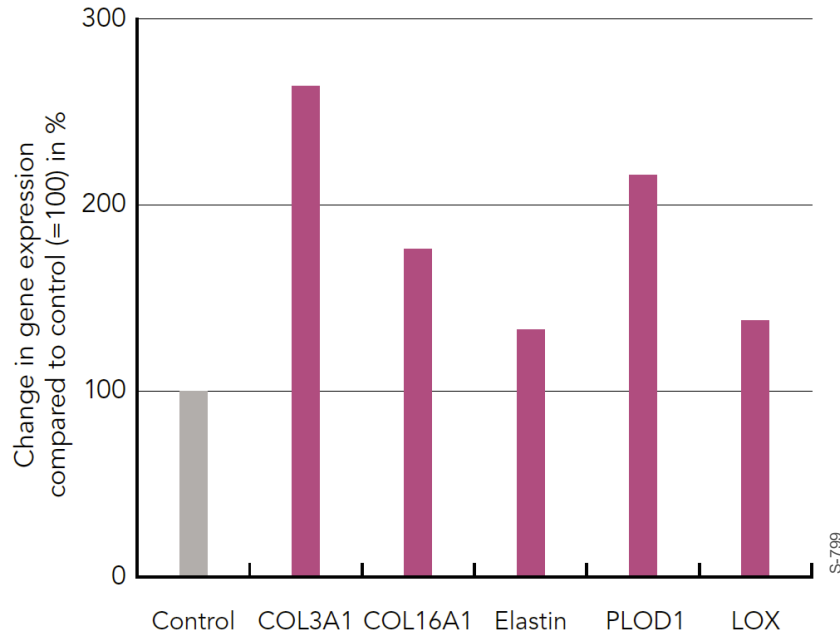


Parameters:

Gene expression of ECM proteins (RT-qPCR).

Stimulation of Extracellular Matrix Genes Through Cell-Cell Communication

- Conditioned medium from untreated MSCs
- Conditioned medium from MSCs treated with 1 % Goji stem cell extract



Increased gene expression of

- collagen 3 and 16*
- elastin*
- PLOD1 (crucial for collagen production)*
- LOX (connects collagen and elastin → stability and elasticity ↑)*

→ PhytoCellTec™ Exosomes improves cell-cell communication between MSCs and fibroblasts.

* No effect via direct treatment of fibroblasts.

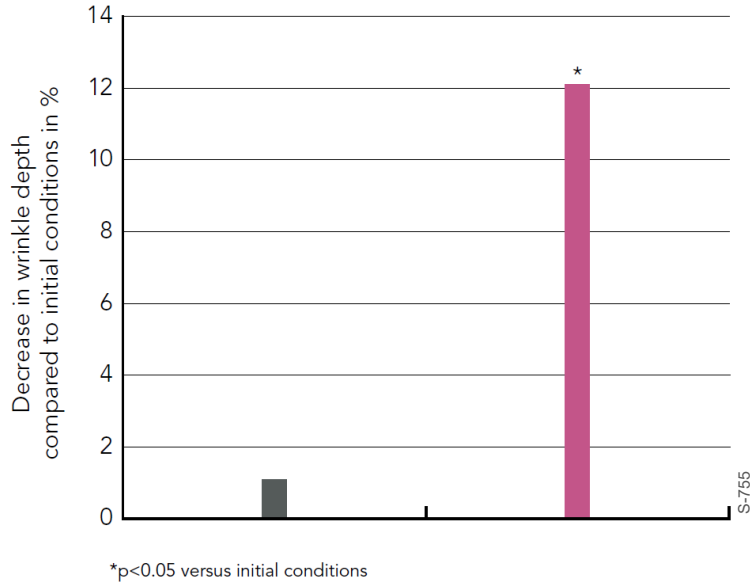


Improvement of Fine Lines and Wrinkles

Volunteers:	23 (f, Caucasian, 41 - 69 y), with signs of photo-aging
Test substance:	Cream with 0.4 % PhytoCellTec™ Exosomes, placebo cream
Application:	Twice daily on one-half of the face, the corresponding placebo on the other half of the face, for 56 days.
Parameter:	Wrinkle depth (PRIMOS lite)

Improvement of Wrinkles

■ Placebo ■ 0.4 % PhytoCellTec™ Exosomes



PhytoCellTec™ Exosomes significantly improved wrinkle depth after 2 months.

Before



After 56 days



Fine wrinkles
appear smoother.

Before



After 56 days



Deeper lines are
visibly reduced.

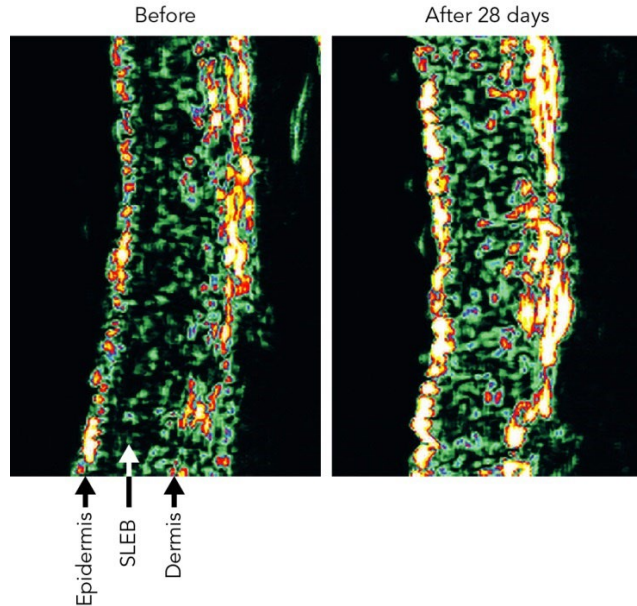


Improvement of Skin Density in Photo-Aged Skin

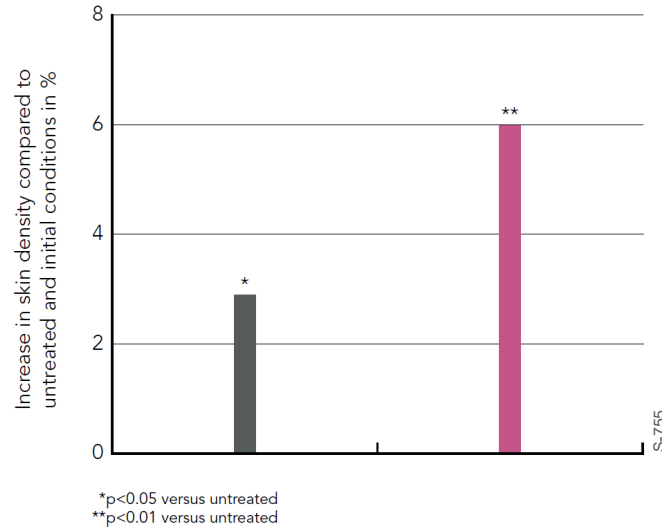
In an intact, youthful dermis, the collagen and elastic fiber structure is dense and yields colorful reflections in ultrasonographic pictures. In photo-aged skin, **disruption of this collagen structure** leads to dark patches (= subepidermal low-echogenic bands, **SLEB**).

Volunteers:	23 (f, Caucasian, 41 - 69 y), with signs of photo-aging
Test substance:	Cream with 0.4 % PhytoCellTec™ Exosomes, placebo cream
Application:	Twice daily on the inner side of one forearm (placebo cream on the inner side of the other forearm) for a period of 28 days
Parameter:	Density (epidermis + dermis) by ultrasonic measurements

Improvement of Skin Density



■ Placebo ■ 0.4% PhytoCellTec™ Exosomes



PhytoCellTec™ Exosomes significantly improved **skin density** and visibly **reduced the SLEB** in photo-aged skin after 1 month.



Improvement of Oval Face Shape

Volunteers:

- 67 (f, Caucasian, 39 - 70 y) with sagging facial skin,
- ✓ split into two groups:
 - ✓ Group 1 applied a cream with 0.4 % PhytoCellTec™ Exosomes
 - ✓ Group 2 applied the corresponding placebo cream

Test substance:

Cream with 0.4 % PhytoCellTec™ Exosomes, placebo cream

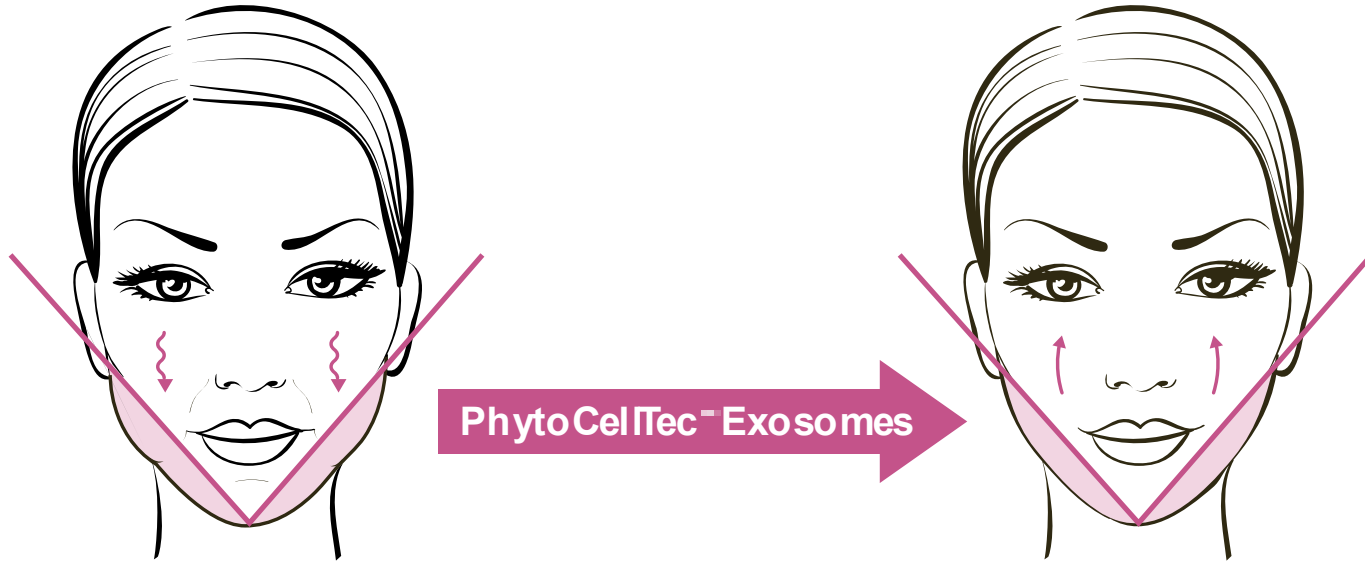
Application:

Face and neck, twice daily for 28 days

Parameter:

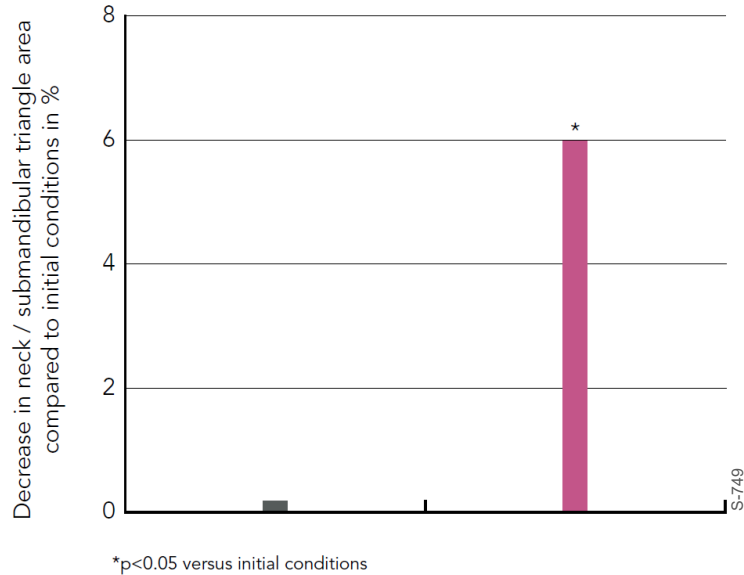
Oval face shape = neck / submandibular triangle size (Visioface)

Oval Face Shape Measurement



Improvement of Oval Face Shape

■ Placebo ■ 0.4 % PhytoCellTec™ Exosomes



PhytoCellTec™ Exosomes significantly improved oval face shape.

Improvement of Oval Face Shape

Before



After
28 days



S-749 / © Mibelle Biochemistry



Breast Lifting Effect

Aging leads to reduced extracellular matrix production,

less elastic and sagging skin

→ lowering of the breast

→ loss of fullness of the breast

Volunteers:

44 (f, 18 - 44 y), split into two groups

Test substance:

Cream with 0.4 % PhytoCellTec™ Exosomes,
placebo cream

Application:

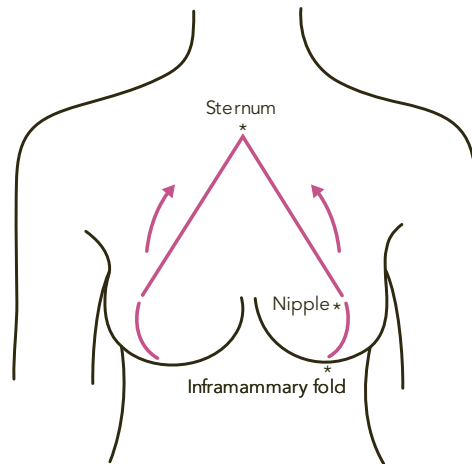
Twice daily for 56 days on the breast area

Parameter:

Bust distances (VECTRA-XT)

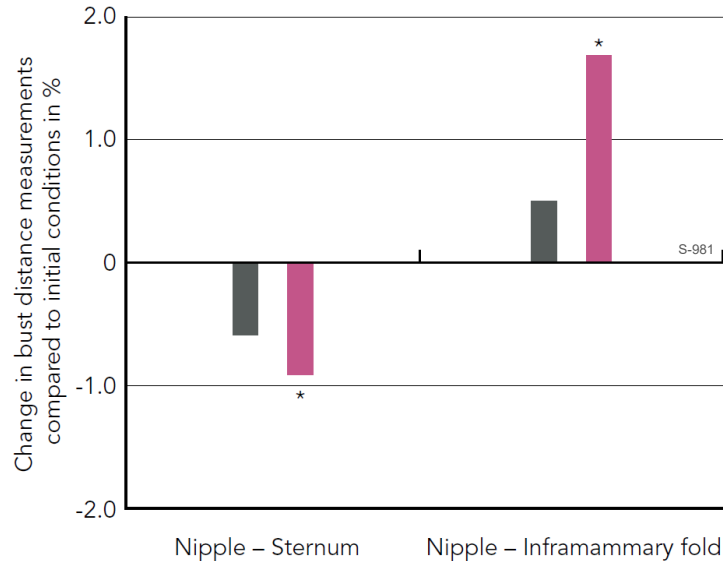
Skin elasticity (Cutometer dual MPA 580)

Breast condition (Regnault ptosis classification)

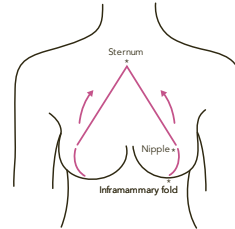


Improvement of Breast Lifting after 2 Months

■ Placebo ■ 0.4 % PhytoCellTec™ Exosomes



*p<0.05 versus initial conditions



Measurement of the bust distances between

- the nipple and the sternum
- the nipple and the inframammary fold

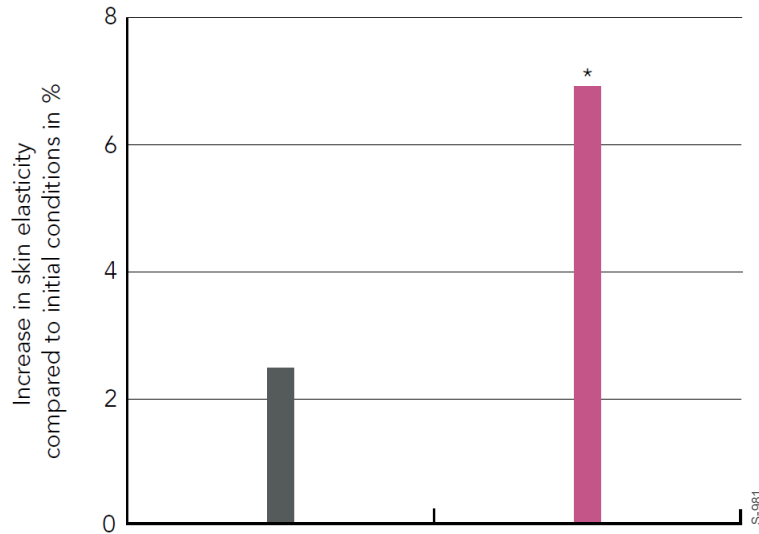
Lifting = reduced nipple-sternum distance
+ increased nipple-inframammary fold distance



PhytoCellTec™ Exosomes significantly improved breast lifting.

Improvement of Skin Elasticity after 2 Months

■ Placebo ■ 0.4 % PhytoCellTec™ Exosomes



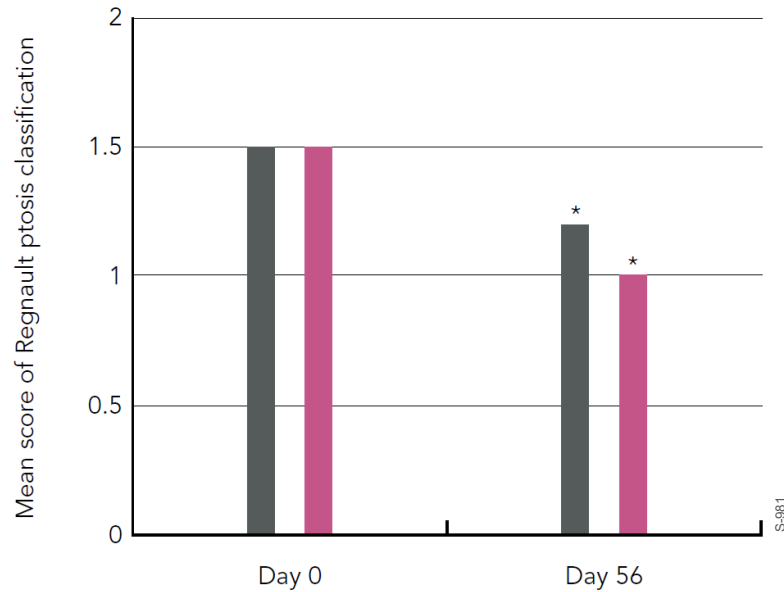
*p<0.05 versus placebo and *p<0.001 versus initial conditions



PhytoCellTec™ Exosomes significantly improved skin elasticity in the breast area.

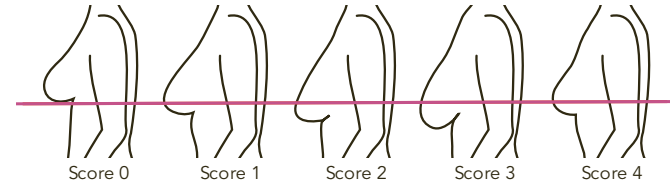
Anti-Sagging Breast Effect after 2 Months

■ Placebo ■ 0.4 % PhytoCellTec™ Exosomes



*p<0.05 versus initial conditions

Regnault ptosis classification



Scheme of the Regnault ptosis classification:
(Score 0) normal, (Score 1) mild sagging, (Score 2) moderate sagging,
(Score 3) severe sagging, (Score 4) pseudoptosis.



PhytoCellTec™ Exosomes significantly improved breast lifting in 54.5% of the volunteers.

PhytoCellTec™ Exosomes



- Doubles the exosomal power
- Increases collagen and elastin expression
- Improves skin density
- Minimizes fine wrinkles and deeper lines
- Tightens facial contours
- Breast lifting effect

PhytoCellTec™ Exosomes Applications



- Exosome cosmetics
- Tightening and contouring serums for the face and neck
- Collagen boosting formulations
- Contouring masks
- Lifting and firming anti-aging formulas
- Breast lifting treatments

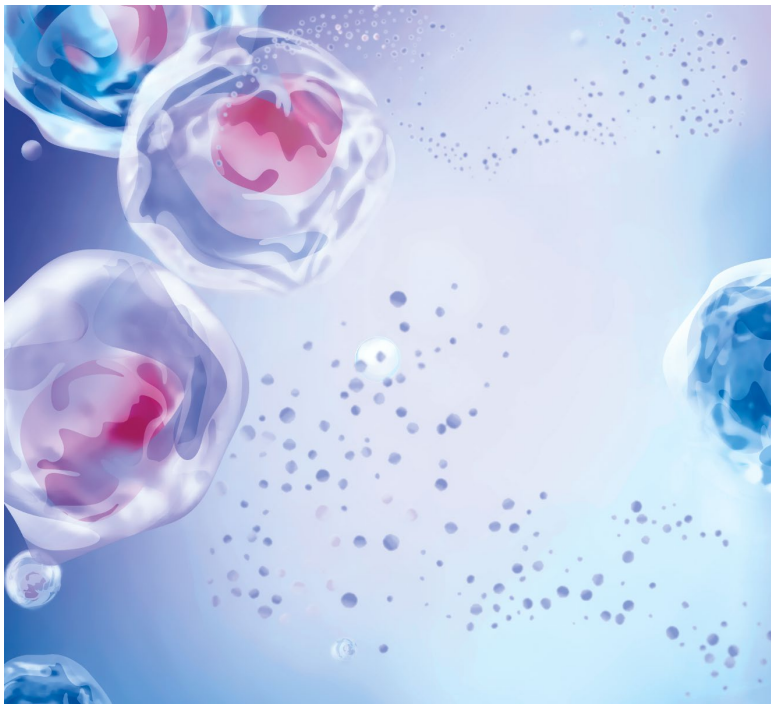
PhytoCellTec™ Exosomes Marketing Benefits



- Dual exosome activity
- Advanced stem cell cosmetics
- Plant stem cells from a superfruit
- Application of exosome signaling
- Proven efficacy on mesenchymal stem cells
- Sustainable production of raw material
- BSB Silver Award - Category Cosmetics – Raw Materials / Actives
- Innovation Zone Best Ingredient Bronze Award Category Actives
- [Product movie](#)
- www.phytocelltec.ch



PhytoCellTec™ Exosomes Marketing Tools



Prototype Formulation

PhytoCellTec™ Exosomes Rejuvenating
Sheet Mask

→ PDF available, samples available, [blog](#)



Website

[Blog post](#)

[Product description](#)

Movies

[Product movie](#)

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