

APIVITA



Ο ΧΗΜΙΚΟΣ ΣΤΗ ΒΙΟΜΗΧΑΝΙΑ ΤΩΝ ΚΑΛΛΥΝΤΙΚΩΝ

ΚΩΝΣΤΑΝΤΙΝΟΣ ΓΑΡΔΙΚΗΣ, R&D DIRECTOR, APIVITA



1. Laboratories

Phytochemistry
Biochemistry

APIVITA



IN-HOUSE LABORATORIES

RESEARCH

Phyto & Bio-labs

PHYTOLAB



IN HOUSE EXTRACTS
CREATION-ASSESSMENT &
PRODUCTION

BIOLAB



IN HOUSE EFFICACY TESTS
ON HUMAN GENOME

Phytolab

PRESSURIZED LIQUID EXTRACTION

Plant extracts



- Only green and edible solvents
- Reproducibility
- Prevention of oxidation, degradation
- Effective extraction (excellent extraction efficacy for secondary metabolites)
- Minimum energy consumption
- 0 waste process

Current situation

APIVITA has created **>100 in-house extracts** over the years

It is currently producing **52 of those extracts** to an extend of **10 tons per year (2021)**

This represents **70%** of the total extracts consumption (2021)



Phytolab
IN HOUSE SOIL ENHANCER PRODUCTION

Waste management



Process

- Mixing Dry Organic Materials (dry plants, wood chips, saw dust) with the Fresh Organic Matter (whole plants, green grass clippings and leaves) (10d)
- Adding Fresh Plant Material, Biocatalytic, Zeolite and Water (5d)
- Adding Organic wastes (plant waste from extraction operations) (10d)
- Adding Organic wastes (plant waste from extraction operations), Biocatalytic, Zeolite, Nitrogen and Water (~3m)

➤ Soil Enhancer Production

- High Quality
- Nitrogen ➡ High Concentration
- Increased soil water holding capacity
- Organic solvents ➡ not detected



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Biolab

Eco-assessment

Eco-toxicity



Ecotoxicology has been defined as, "the branch of toxicology concerned with the study of toxic effects, caused by natural or synthetic pollutants, to the constituents of ecosystems, animal (including human), vegetable and microbial, in an integral context"

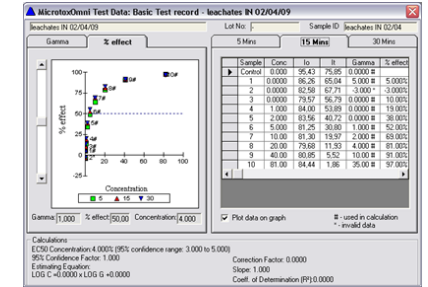
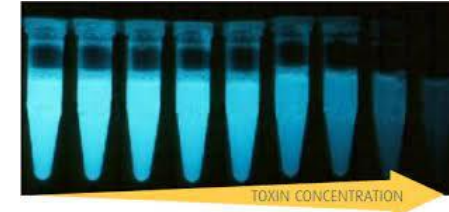
Way to evaluate formulas and ingredients that are environmentally persistent, bioactive, and have the potential for bioaccumulation.

The objective is to evaluate and estimate hazards as a result of personal care products released into the aquatic environment.

- Standardized → different organizations (ISO, OCDE, USEPA, ASTM, etc.)
- Toxicity → acute and chronic.

Vibrio fischeri

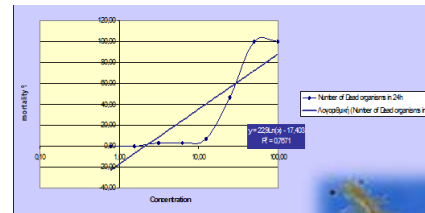
EC50 is the concentration that causes adverse effects in 50% of the test organisms



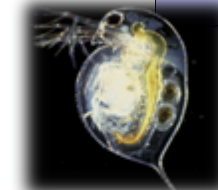
LC50 is the acute toxicity, the lethal concentration at which 50% of the test organism dies within the test-specified time. The test may start with eggs, embryos, or juveniles and last from 24 hours to 96 hours [citation needed]..

Seawater

Artemia franchiscana



Freshwater Daphnia magna





2D cell cultures and 3D skin models

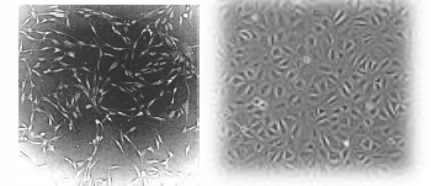
An alternative to animal testing. These cell cultures include not only keratinocytes but also other skin cell types, such as fibroblasts, which are predominantly in the dermal layer of the skin, and certain immune cells and even melanocytes.

In-house applications:

- **2D assays**
- ✓ **Safety assessments:** Cell viability, proliferation and metabolism (MTT assay)
- ✓ **Efficacy assessments:** Transcriptomic analysis, protein quantification, wound healing, epigenetics
- **3D assays**
- ✓ **Skin sensitization: 3d reconstructed human skin Epi-skin**
- ✓ **Eye irritation : EpiOcular**

In Vitro cell models

2D culture cells



Normal Human primary Dermal Fibroblasts (NHDF)

Normal Human primary Epidermal Keratinocytes (NHEK)

3D skin models

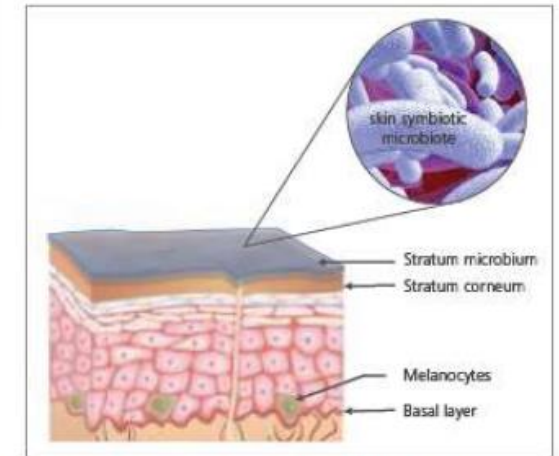
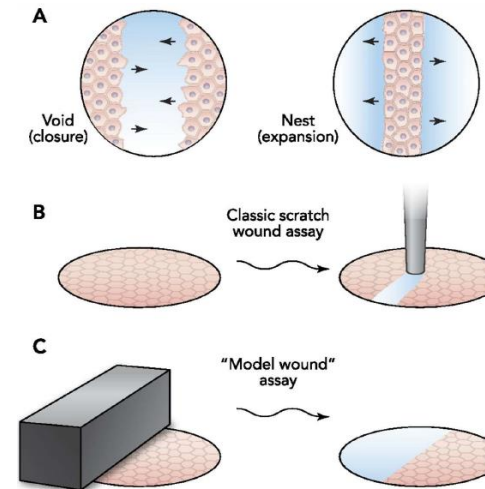
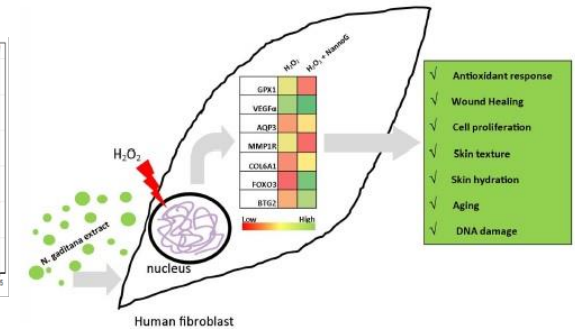
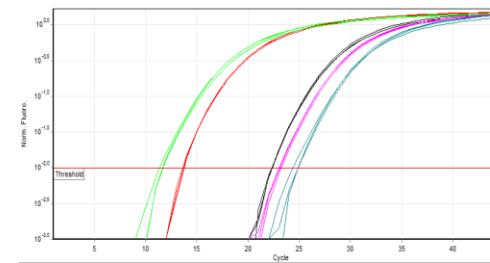


Biolab



Efficacy Assessments-Proof of efficacy

- Transcriptomic profiling method: detection of expression **200 genes** involved in skin related processes (**APIGENES** platform)
- **Epigenetics** : methylation process, detection of miRNAs related to aging
- **Protein Biochemistry** assays to further validate the genomic data in the **post-transcriptional level**
- **Metagenomics** : target skin microbiome - effect on stratum microbium
- **In vitro migration** assays in 2D cell cultures to study the **wound healing** effect of tested extracts



Fitz-Gibbon et al., Journal of Investigating Dermatology, 2013

IN HOUSE EFFICACY TESTS- A CASE STUDY

Biolab

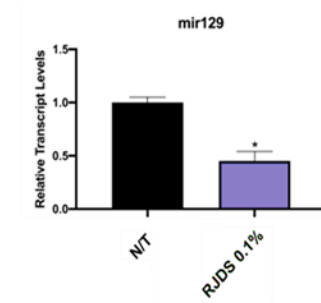
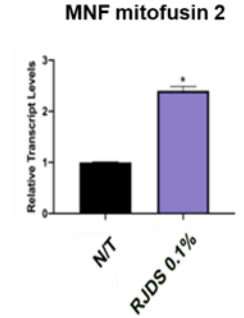
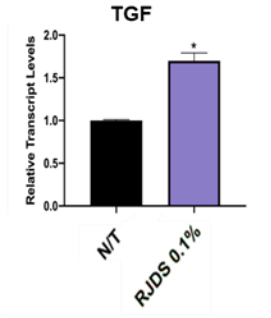
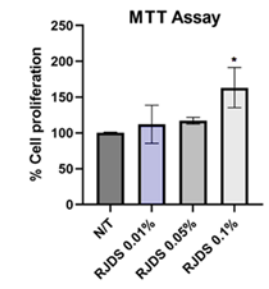
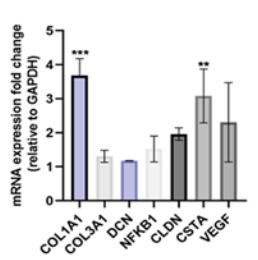
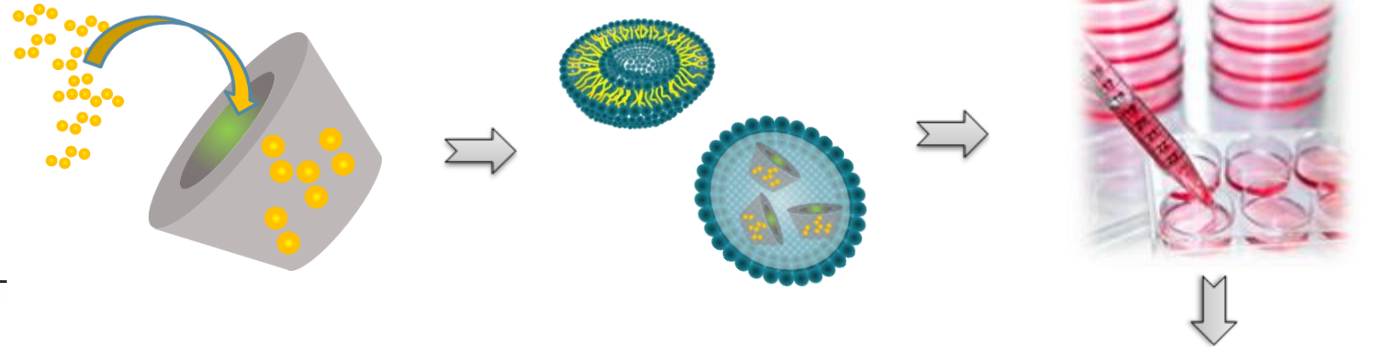


Royal Jelly Controlled Release

Protects fibroblasts and ECM by enhancing **collagen production, cell contractility** (Col1A1, CSTA)

Promotes **cell proliferation, cell viability and metabolism** in human fibroblasts (MTT assay, TGF, MNF2)

Protects fibroblasts from **aging** process (mir129- Epigenetic biomarker)



2. Skin Delivery Systems

Adding value to bioresources
Targeted delivery

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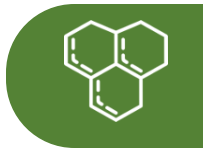
INNOVATION

Representative classes of plant-derived bioactive compounds

Challenges in the Use of Natural Products



Solubility



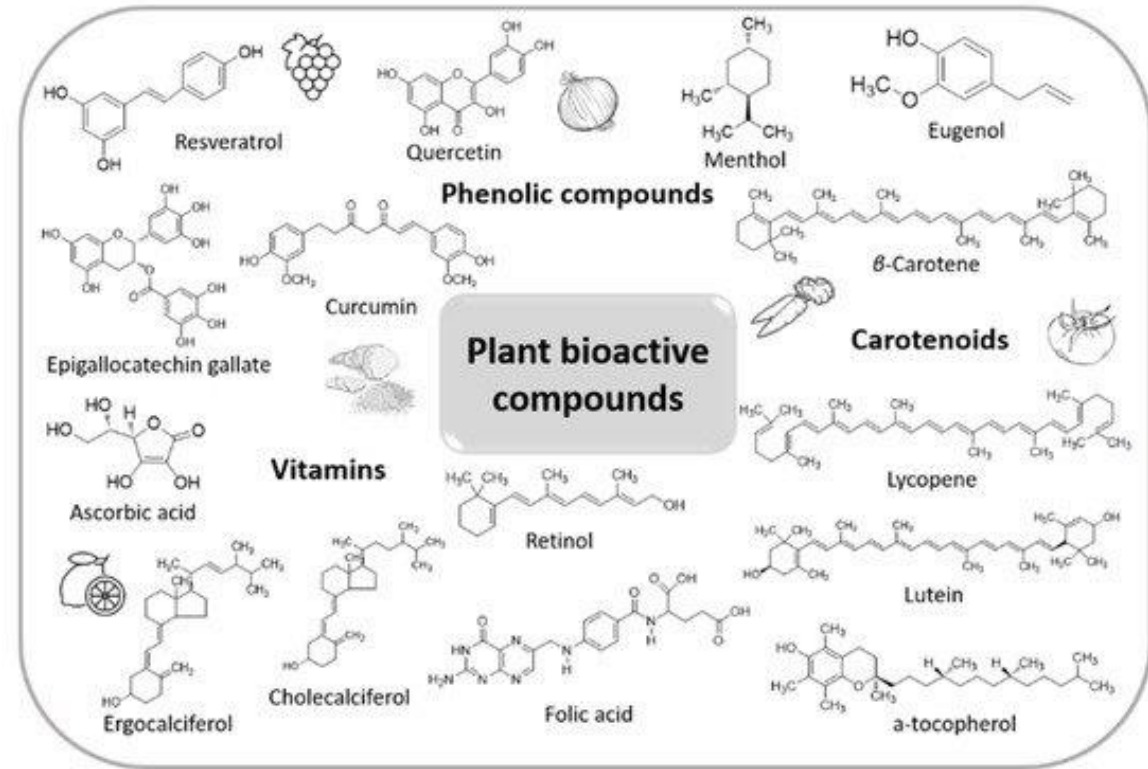
Stability



Bioavailability



Release rate

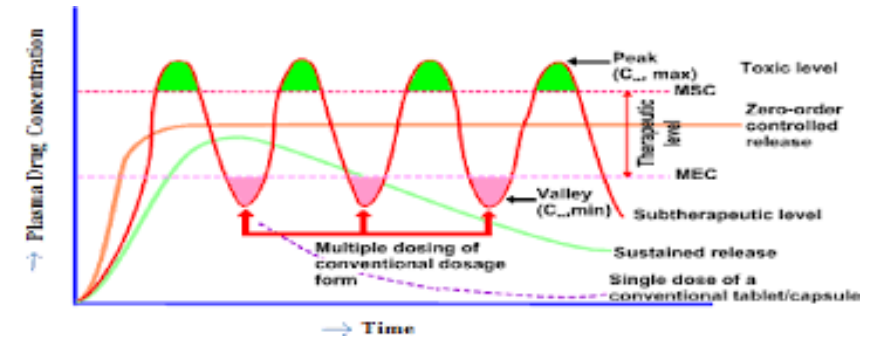
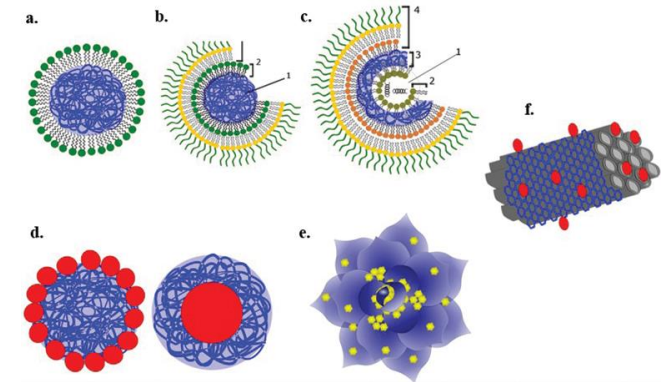
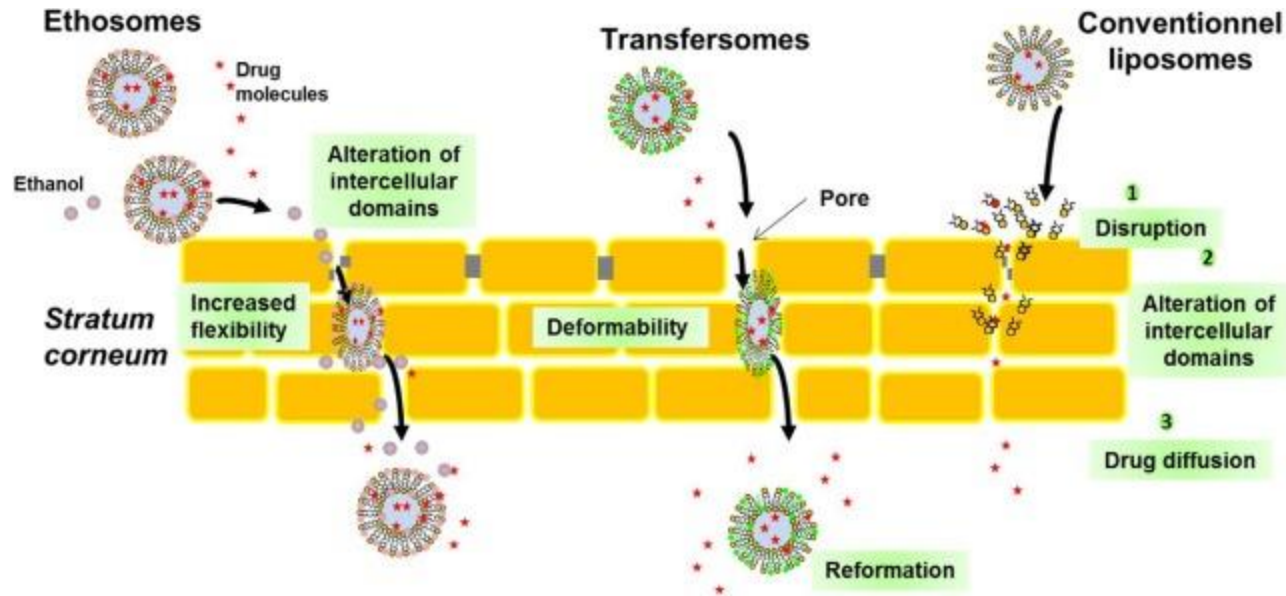


INNOVATION

Combinatorial systems

Skin penetration/Release rate

SKIN DELIVERY SYSTEMS

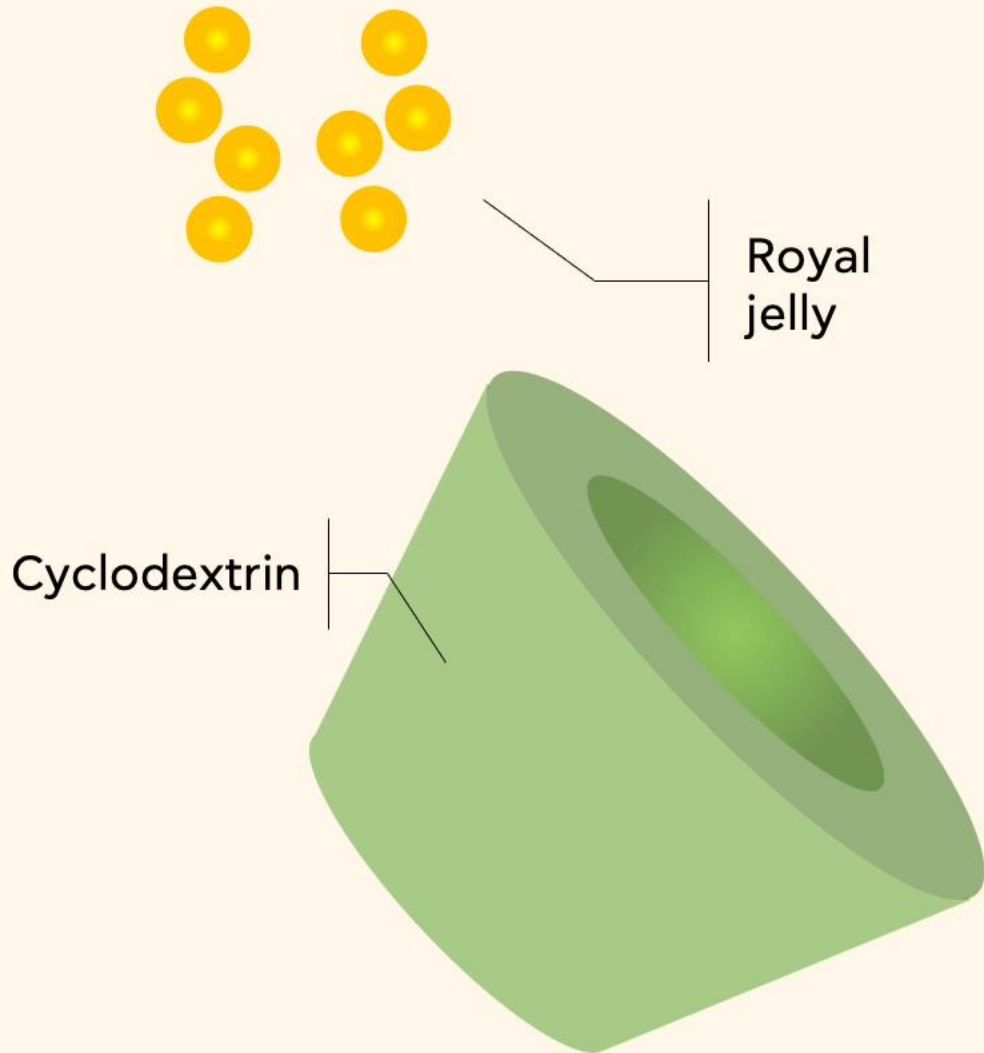


M. Sala, R. Diab, A. Elaissari, H. Fessi, Lipid nanocarriers as skin drug delivery systems: Properties, mechanisms of skin interactions and medical applications, International Journal of Pharmaceutics, Volume 535, Issues 1–2, 2018, Pages 1–17
 Madni, A. et al., 2017, 'Hybrid Nano-carriers for Potential Drug Delivery', in S. Maiti, K. K. Sen (eds.), Advanced Technology for Delivering Therapeutics, IntechOpen, London. 10.5772/66466.

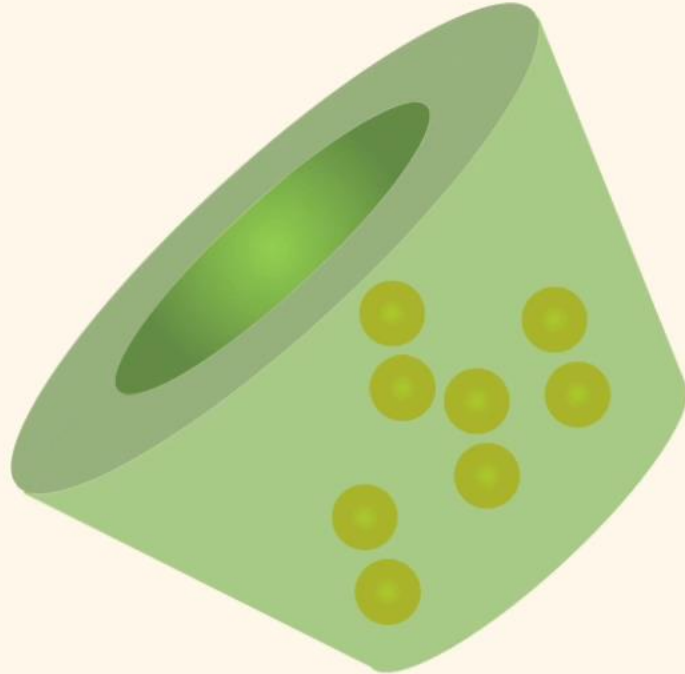
HYBRID NATURAL CARRIER

PHASE 1:

Royal jelly is encapsulated inside a cyclodextrin for maximum protection



HYBRID NATURAL CARRIER

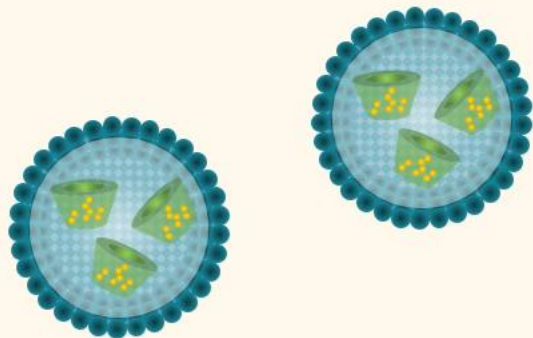


PHASE 2:

The cyclodextrin containing the royal jelly is in turn encapsulated in the core of a liposome

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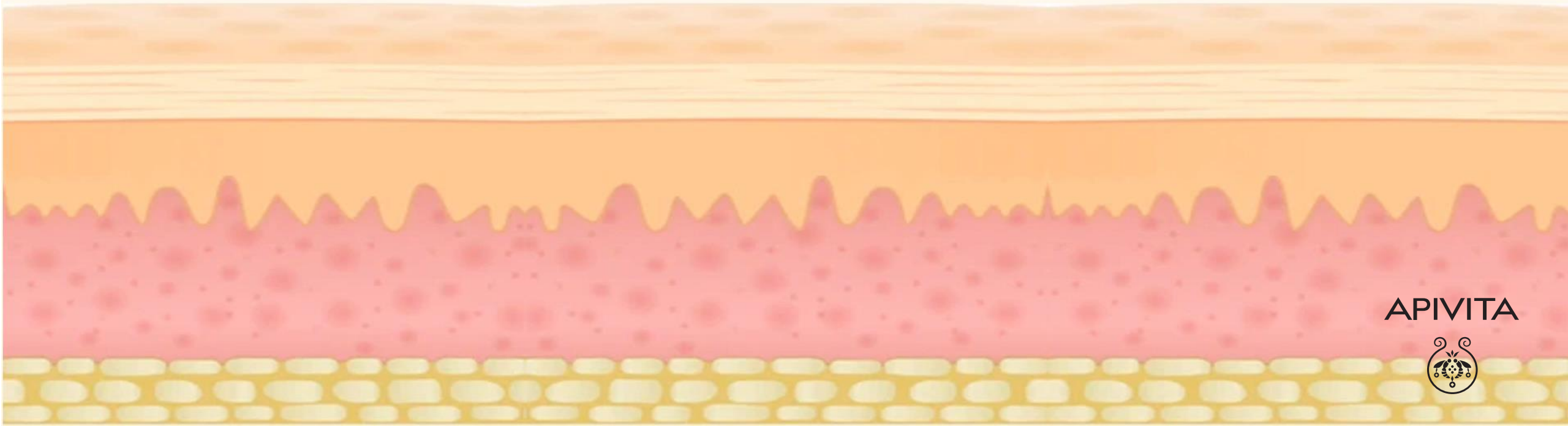


HYBRID NATURAL CARRIER

PHASE 3:

Deep & time controlled release

Bioactivity of royal jelly offered to skin in a time-controlled way for maximum absorption and efficacy



APIVITA



3. COLLABORATIVE RESEARCH

Funded research programs

APIVITA



Academic ecosystem targeting innovation in sustainability

Innovation Greentech expertise

RESEARCH PROGRAMS

RESEARCH PROGRAMS



Long-term research collaborations

Four “2020 horizon” international research programs

- ✓ 16 accomplished Research Programs
- ✓ 16 M euro budget
- ✓ 12 Established lab methods
- ✓ 7 products on market
- ✓ 2 innovative products pending
- ✓ 78 collaborating institutes/universities

UNIVERSITIES PARTNERSHIPS



Prestigious partnerships with international Academia

Active collaborations with 25 countries around the world

4. FORMULATION

Innovative effective final products

APIVITA



Development

PRODUCT DEVELOPMENT PHILOSOPHY

We provide the most effective solution for skin's well being following all **intrinsic** and **restoring** mechanisms

- Unlock all mechanisms for skin's best function
- Maximize the inner possibilities for flawless skin

Formulation is driven by **sophisticated simplicity**

- All included ingredients have a very special synergistic mission to deliver in the formula

Health and beauty are indispensably connected

- Cutting edge technologies to unblock and enhance skin's role to health

We seek optimum balance between **maximum results** and **green+clean** formulations

- Effective ingredients that have been selected according to their sustainability and clean profile



CLEAN FORMULATION CHARTER

NATURAL



SAFE



EFFECTIVE



SUSTAINABLE



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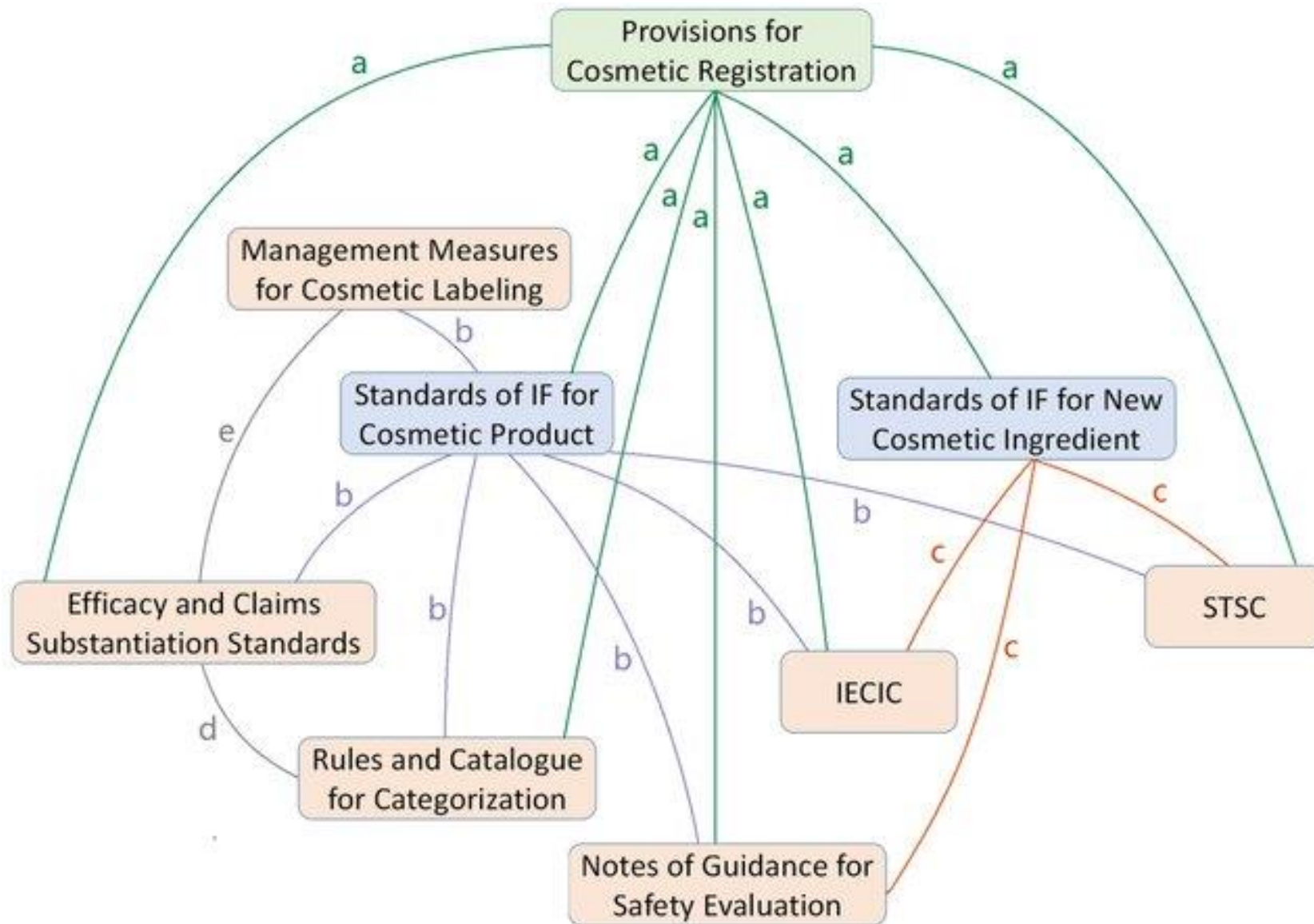


5. REGULATORY

Legal consolidation of portfolio

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6. SCIENCE COMMUNICATION

Dissemination of scientific expertise

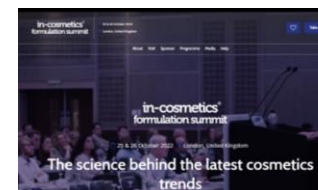
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A tool for brand marketing


Recognition Scientific communication

- Keynote speakers in global leading scientific conferences
- Publications in peer-reviewed scientific journals with impact factor
- TV, radio, newspaper, social media communication of APIVITA scientific excellence linked with product launches
- Scientific presentation of innovation during product launches events
- Organization of innovation days, workshops, summer schools aiming at the brand's scientific image reinforcement



Review

Innovative Delivery Systems Loaded with Plant Bioactive Ingredients: Formulation Approaches and Applications

Anastasia Kyriakoudi ¹, Eleni Spanidi ², Ioannis Mourtzinis ¹  and Konstantinos Gardikis ^{2,*}

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Article




Honey Extracts Exhibit Cytoprotective Properties against UVB-Induced Photodamage in Human Experimental Skin Models

Athanasios Karapetsas ¹, Georgia-Persephoni Voulgaridou ¹, Dimitra Iliadi ¹, Ilias Tsochantaridis ¹, Panagiota Michail ¹, Spyridon Kynigopoulos ², Maria Lam Maria-Ioanna Stavropoulou ³, Konstantina Stathopoulou ³, Sofia Karabournioti ⁴, Nektarios Aligiannis ⁵, Konstantinos Gardikis ⁵, Alex Galanis ¹, Mihalis I. Panay; Aglaia Pappa ^{1,*}



Article

Propolis Extracts Inhibit UV-Induced Photodamage in Human Experimental In Vitro Skin Models

Athanasios Karapetsas ^{1,†}, Georgia-Persephoni Voulgaridou ^{1,†} , Manolis Konialis ¹, Ilias Tsochantaridis ¹, Spyridon Kynigopoulos ², Maria Lambropoulou ² , Maria-Ioanna Stavropoulou ³, Konstantina Stathopoulou ³, Nektarios Aligiannis ³, Petros Bozidis ⁴, Anna Goussia ⁴, Konstantinos Gardikis ⁵, Mihalis I. Panayiotidis ⁶  and Aglaia Pappa ^{1,*}

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The Journal of Supercritical Fluids

Volume 146, April 2019, Pages 159-164



Supercritical CO₂ extraction of *Salvia fruticosa*

Daphne Kavoura ³, Konstantina Kyriakopoulou ³, Georgios Papaefstathiou ³, Eleni Spanidi ⁴, Konstantinos Gardikis ⁴, Vasiliki Louli ³ , Nektarios Aligiannis ⁵, Magdalini Krokida ⁶, Konstantinos Magoulas ³

Short Commentary

Photoprotective Properties of Honey Extracts and their Correlation with the Metabolomic Content








Konstantinos Gardikis

APIVITA SA, Industrial Park of Markopoulo Mesogaia, 19003 Markopoulo Attiki, Athens, Greece



Article

A New Controlled Release System for Propolis Polyphenols and Its Biochemical Activity for Skin Applications

Eleni Spanidi ¹, Athanasios Karapetsas ², Georgia-Persephoni Voulgaridou ² , Sophia Letsiou ¹ , Nektarios Aligiannis ³ , Ilias Tsochantaridis ² , Spyridon Kynigopoulos ⁴, Maria Lambropoulou ⁴ , Ioannis Mourtzinis ⁵ , Aglaia Pappa ²  and Konstantinos Gardikis ^{1,*}

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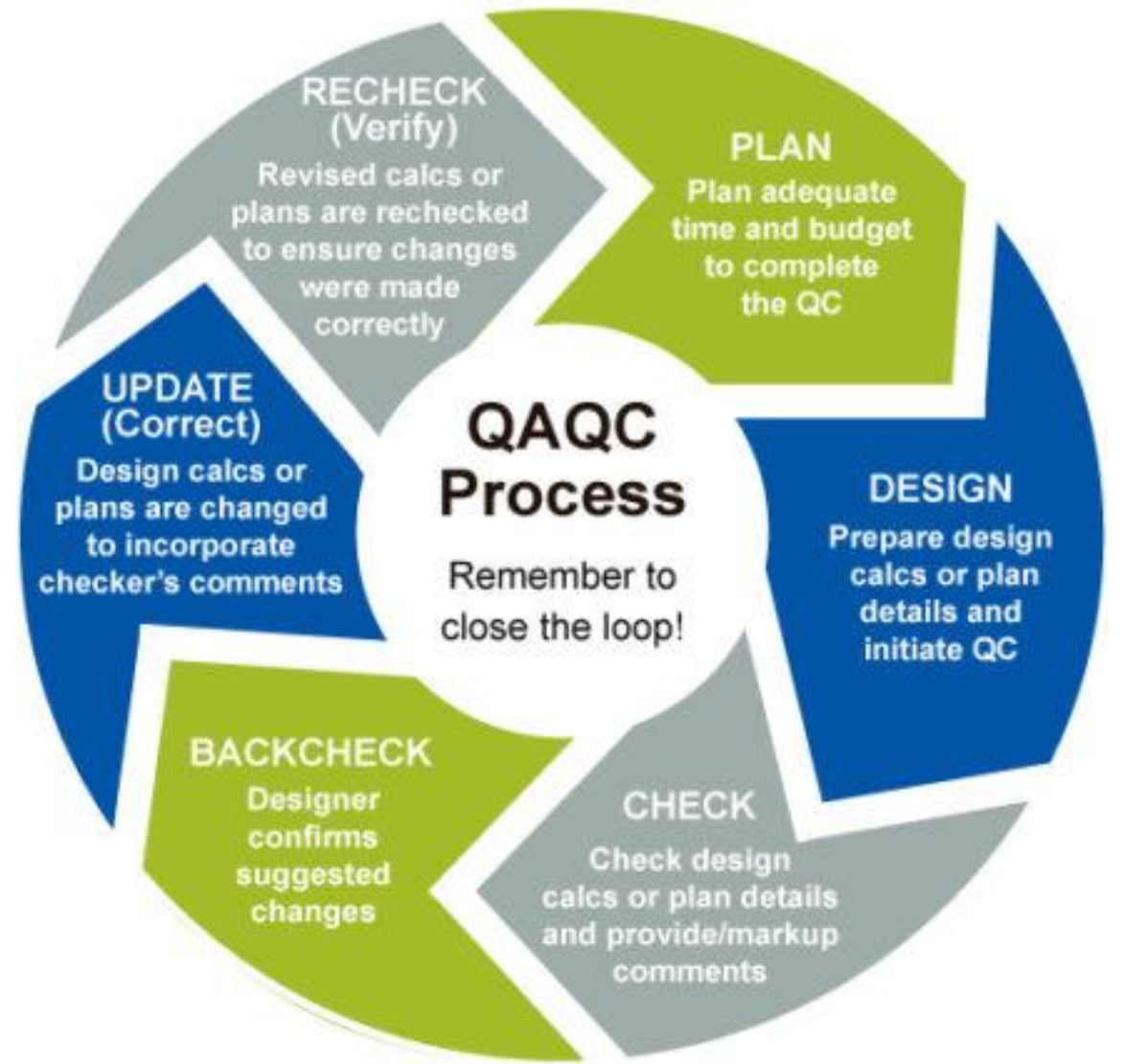
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FULL PAPER



Photometric Analysis of Propolis from the Island of Samothraki, Greece. The Discovery of Red Propolis

Andros Papachristoforou, ^{a,b} Evgenia Louvoula, ^c George Menexes, ^d Konstantinos Gardikis, ^e and Ioannis Mourtzinis ^f



Essentials Of A Cosmetics Business Plan

APIVITA



ΕΥΧΑΡΙΣΤΩ!