EPISKIN World leader in tissue engineering

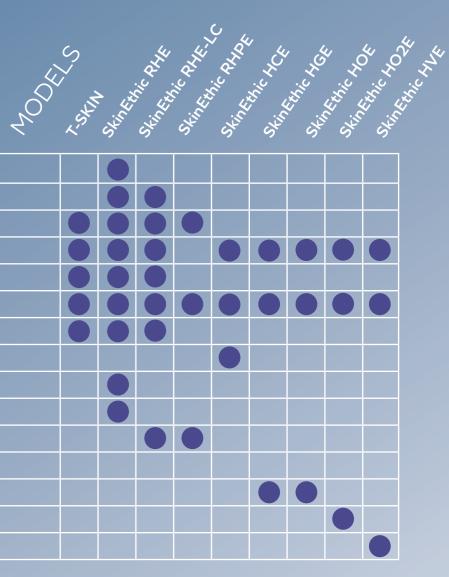
FOR IN VITRO TESTING

APPLICATIONS

Skin irritation	
Skin corrosion	
UV exposure	
Bacterial adhesion	
DNA Damage	
Omics	
Permeability	
Eye irritation	
Medical Devices	
Microbiome	
Skin immune response	
Pigmentation/Depigmentation	
Oral & Gingival care	
Oesophageal Irritation	
Vaginal irritation	

SUMMARY

T-SKIN[™] / Reconstructed Full-Thickness N SkinEthic[™] RHE / Reconstructed Human SkinEthic[™] RHE-LC / Human Epidermal N SkinEthic[™] RHPE / Reconstructed Human SkinEthic[™] HCE / Human Corneal Epithel SkinEthic[™] HOE / Human Oral Epithelium SkinEthic[™] HGE / Human Gingival Epithel SkinEthic[™] HO2E / Human Oesophageal SkinEthic[™] HVE / Human Vaginal Epithelium



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n Pigmented Epidermis	11
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lium	17
Epithelium	19
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EPISKIN: your reliable and trustworthy partner for your benefits

We innovate

since 1992 for a better science in producing advanced, relevant and robust in vitro 3D models allowing our users to predict human response. Very strict and unique quality controls help us to supply unmatched quality models. Our innovation, your science.

We deliver

Innovation

Expertise

worldwide, serving all customers with our 3 partners in India, Korea and Japan and with our 2 subsidiaries in Brazil and China and our headquarters in France. You order, we deliver.

ADVANCED 3D MODELS FOR A BETTER, MORE PREDICTIVE AND MORE ETHICAL SCIENCE



all our expertise, strengths and skills to be the world leader in tissue engineering for in vitro testing and research. No compromise. Our one and only core business is supplying advanced and relevant tissues to the scientific community and partnering with added value testing laboratories. Our commitment, your satisfaction.

NAME

DESCRIPTION

T-Skin[™] is an in vitro reconstructed skin which consists of a dermal equivalent with human fibroblasts overlaid by a stratified, well differentiated epidermis derived from normal human keratinocytes cultured on an inert polycarbonate filter.

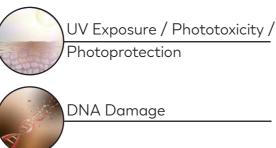
This model exists at different stages of maturity.

Reconstructed Human Full Thickness Model

APPLICATIONS

FORMAT

6-well plate





Omics

Permeability

SPECIFIC MARKERS

Differentiation markers:

Filaggrin Involucrin

- Loricrin
- Cytokeratin 10
- Transglutaminase-1

Dermal-Epidermal junction markers

- Collagen IV, VII, XII
- Laminin V
- Perlecan **BP** antigen

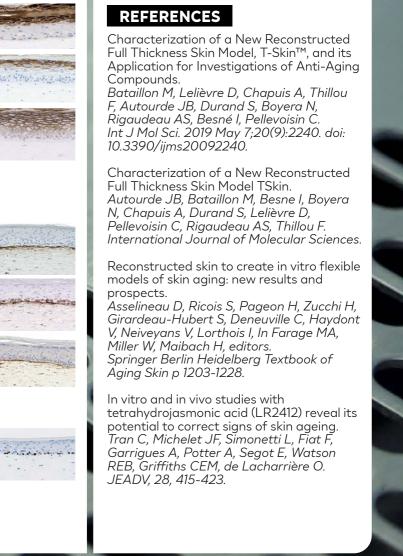
Proliferation markers

Ki67



Reconstructed Human Full Thickness Model

T-Skin[™] / Reconstructed Human Full Thickness Skin Model



RHE SkinEthic

hold of a first first as a familia the

Epidermis

NAME

DESCRIPTION

SkinEthic[™] RHE is an *in vitro* reconstructed human epidermis from normal human keratinocytes cultured on an inert polycarbonate filter at the air-liquid interface. It is histologically similar to the in vivo human epidermis.

Our strong believe in Science and our continuous improvement with ISO 9001 certification push us to keep improving the production process of our model: From cell extraction to reconstruction with chemically defined biocomponents and medium.

Every single biocomponent of each step of our production is clearly defined and their traceability is guaranteed. The process is then more secured, allowing to deliver a SkinEthic[™] RHE model more reproducible, robust and reliable than ever.

Different maturities and surfaces are available.

SPECIFIC MARKERS

Differentiation markers:

- Filaggrin
- Involucrin Loricrin
- Transglutaminase-1 Keratin 10 Keratin 5 CD44

AVER OF CONTRACTOR

Presence of different epidermal classes of lipids comprising ceramides

Dermal-Epidermal junction markers

- Type IV collagen
- Laminin V Alpha6Beta4-integrin BP antigen



Proliferation markers Ki67

Cell Migration Model ΩМ **S**kin**E**thic

are epidermal models on an innovative insert. Epidermal or mucosal reconstructed on this insert are histologically similar to in vivo human epidermis and mucosa.

Reconstructed Human Epidermis

APPLICATIONS



- UV Exposure / Phototoxicity / Photoprotection
- DNA Damage



Omics

Permeability

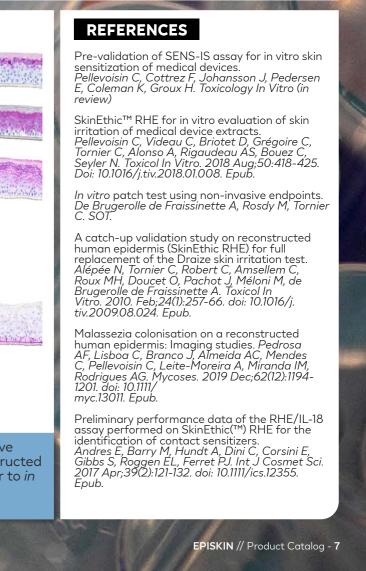
Medical Device

FORMAT



Reconstructed Human

SkinEthic[™] RHE / Reconstructed Human Epidermis



RHE-LC



NAME

Cells

DESCRIPTION

The SkinEthic[™] RHE-LC model is a standard epidermal model in which Langerhans cells progenitors have been integrated. During the tissue reconstruction, these immature cells have differentiated into antigen-presenting Langerhans cells expressing the specific marker CD207 (langerin). They are mostly located and evenly spread within the supra-basal epidermal layer. This model is therefore expected to be a useful tool for skin immune response studies.

Reconstructed Human Epidermis with Langerhans Cells

APPLICATIONS



UV Exposure / Phototoxicity /
Photoprotection

Bacterial adhesion

Permeability

Skin immune response

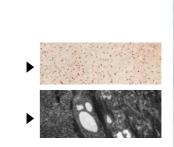


SPECIFIC MARKERS

Langerhans cells markers:

CD207 Langerin Birbeck granules

CD1 CD80 CD86 CCR7



Differentiation markers:

Filaggrin Involucrin Loricrin Keratin 10 Keratin 5

Presence of different epidermal classes of lipids comprising ceramides

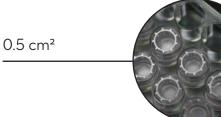
Dermal-Epidermal junction markers

Type IV collagen

Laminin V ► Alpha6Beta4-integrin BP antigen

Proliferation markers Ki67





Reconstructed Human Epidermis with Langerhans Cells

SkinEthic[™] RHE-LC / Human Epidermal Model Langerhans

REFERENCES

Adding the immune component in reconstructed human skin and eye epithelia models

moaels. Ancelin N, Benas D, Brusq JM, Douillard G, Ligouis M, Meloni M, Mondoulet L, Ovigne JM, Pellevoisin C, Rigaudeau AS, Sahuc F, Segaud V. Poster Asiatox

Cellular mechanistic investigation on antigen delivery by Viaskin® patchfor epicutaneous immunotherapy with reconstructed human epidermis including Langerhans cells (SkinEthicTM RHE-LC). Dhelf V, Dioszeghy V, Ligouis M, Mondoulet L, Pellevoisin C, Sahuc F. Poster

RHPE

Reconstructed Human Pigmented Epidermis

NAME

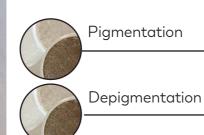
DESCRIPTION

The SkinEthic™ RHPE model is composed of normal human keratinocytes cultivated in the presence of melanocytes of phototype II, IV or VI, localized in the basal layer.

of human skin.

Reconstructed Human Pigmented Epidermis

APPLICATIONS



UV Exposure / Phototoxicity / Photoprotection

Omics



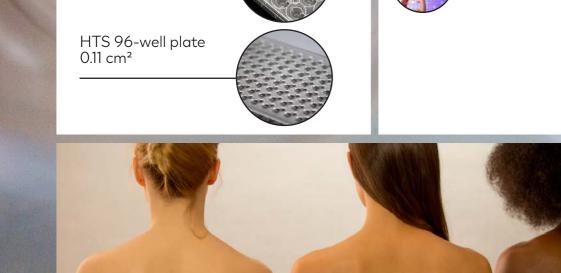
SPECIFIC MARKERS

Melanocytes localized in the basal cell layer interspersed with basal cell keratinocytes.



Melanin distribution in the basal layer

Ki 67



10

FORMAT

0.5 cm²

4 cm²

0.33 cm²

HTS 24-well plate

SkinEthic[™] RHPE / Reconstructed Human Pigmented Epidermis

The different tanning degrees of these constructs correspond macroscopically to 3 different phototypes



REFERENCES

Skin lightening effect of natural extracts coming from Senegal botanical biodiversity. Baillet-Guffroy A, El Khoury R, Lteif R, Michael-Jubeli R, Salameh D, Tfayli A, Zeitoun H.

International Journal of Dermatology.

Reconstructed Human Pigmented Epidermis (RHPE): an in vitro model for the evaluation of melanogenesis. Sahuc F. SOFW Magazine.

Sepicalm VG, a new skin lightening enable to modulate melanogenesis-related genes and to prevent UV-induced pigmentation thanks to its soothing properties. Dumont S, Khaiat A, Puginier M, Stoltz C, Garcia C. SEPPIC

Melanocyte containing human organotypic epidermis as a model to evaluate toxicity of melanin binding substances. Straube F, Junker U, Kretz S, Wolf A. Society of Toxicology, USA.

HCE SkinEthic

Human



DESCRIPTION

The SkinEthic[™] HCE model is composed of transformed human corneal keratinocytes cultivated on an inert polycarbonate filter at the air liquid interface in a chemically defined medium.

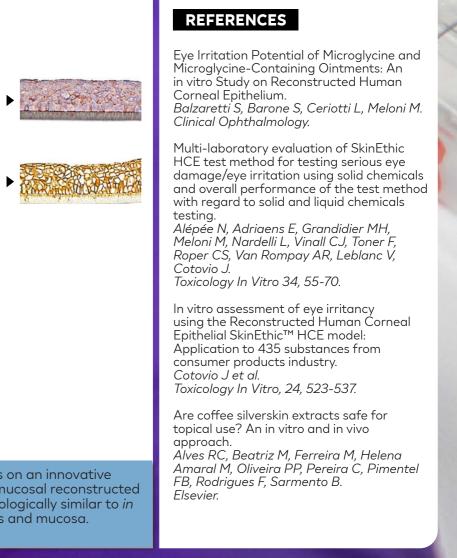
The reconstructed tissue forms a stratified and well organized epithelium which is structurally, morphologically and functionnally similar to the human cornea with presence of basal, wing and mucus production cells.

SPECIFIC MARKERS

Differentiation markers:

Keratin

- CD44
 - Hemidesmosomes





are epidermal models on an innovative insert. Epidermal or mucosal reconstructed on this insert are histologically similar to in vivo human epidermis and mucosa.

Human **Corneal Epithelium**

APPLICATIONS

Ocular Irritation



Bacterial Adhesion

- Eye Irritation Test (EIT) Method validated by EURL-ECVAM and under OECD TG 492 for identification of chemicals not requiring classification for eye hazard (UN GHS)
- ▶ EIT implemented in the draft OECD GD for serious damage and eye irritation
- Corneal differential display i.e. mucin production

12



4 cm²

FORMAT

0.5 cm²



Omics



Corneal Epithelium

SkinEthic[™] HCE / Human Corneal Epithelium

Skin**E**thic

Human



SkinEthic[™] HGE / Human Gingival Epithelium

DESCRIPTION

The SkinEthic[™] HGE model is composed of normal human gingival cells cultivated on an inert polycarbonate filter at the air liquid interface in a chemically defined medium.

human gum.

Human

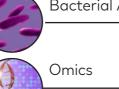
Gingival Epithelium

FORMAT



APPLICATIONS

Oral Care



Bacterial Adhesion

SPECIFIC MARKERS

Differentiation markers:

- Filaggrin
- Keratin 13 Ki67 CK10







Gingival Epithelium

This model is histologically similar to the outer cell layers of the





REFERENCES

Evaluation of an oral care product safety Evaluation of an oral care product safety screening program utilizing the in vitro SkinEthic Human Gingival Epithelium (RHG) and Oral Buccal (RHO) models. Wurzburger L, Kazmi P, Re T, Alonso A, Bertino B, Barnes N, de Brugerolle de Fraissinette A, Hilberer A, Raabe H, Wilt N, Srinivasan V. SOT.

In Vitro Toxicity evaluation of toothpastes using reconstructed human oral and gingival mucosa models. Van de Vannet B, De Wever B, Bottenberg P, Cappadoro M. Presented at the Society of Toxicology, New Orleans, USA.

Characterization of human oral and gingival mucosal models: a histological characterization and applications in toxicity testing.

Vande Vannet B and Hanssens JL, Free University of Brussels. 3rd International SkinEthic Workshop Nice, France.

SkinEthic

Human

NAME

DESCRIPTION

The SkinEthic[™] HOE model is composed of TR146 cells (derived from a squamous cell carcinoma of the buccal mucosa) cultivated on an inert polycarbonate filter at the air liquid interface in a chemically defined medium.

This model forms an epithelial tissue devoid of stratum corneum, resembling histologically to the mucosa of the oral cavity.

Human **Oral Epithelium**

FORMAT

0.5 cm²

HTS 24-well plate 0.33 cm²



APPLICATIONS

Oral Care Bacterial Adhesion



Omics

SPECIFIC MARKERS

Differentiation markers:

Keratin 6



CD44 Ki67





Oral Epithelium

SkinEthic[™] HOE / Human Oral Epithelium



REFERENCES

Evaluation of an oral care product safety Evaluation of an oral care product safety screening program utilizing the in vitro SkinEthic Human Gingival Epithelium (RHG) and Oral Buccal (RHO) models. Wurzburger L, Kazmi P, Re T, Alonso A, Bertino B, Barnes N, de Brugerolle de Fraissinette A, Hilberer A, Raabe H, Wilt N, Srinivasan V. SOT.

A Biphasic Innate Immune MAPK Response Discriminates between the Yeast and Hyphal Forms of Candida albicans in Epithelial Cells. Moyes DL et al. Cell Host & Microbe, 8, 225-235.

Quantitative expression of the Candida albicans secreted aspartyl proteinase gene family in human oral and vaginal candidiasis. Naglik JR et al. Microbiology, 154, 3266-3280.

Phenotypic screening, transcriptional profiling, and comparative genomic analysis of an invasive and non-invasive strain of Candida albicans. Thewes S, Moran GP, Magee BB, Schaller Μ Śullivan DJ, Hube B. BMC Microbiol. 24;8:187.

HO7F

Human

NAME

DESCRIPTION

The SkinEthic[™] H02E model is a human oesophageal epithelium composed of immortalized cell line Kyse 510, cultivated on an inert polycarbonate filter at the air liquid interface in a chemically defined medium.

oesophagus.

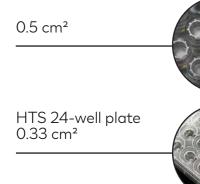
SPECIFIC MARKERS

Differentiation markers:

- Keratin 6
- Keratin 13



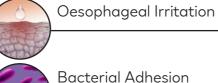
FORMAT



HTS 96-well plate 0.11 cm²



Human



Oesophageal Epithelium

Bacterial Adhesion



Oesophageal Epithelium

SkinEthic[™] HO2E / Human Oesophageal Epithelium

This model forms an epithelial tissue devoid of stratum corneum, resembling histologically to he outer cell layers of the human

REFERENCES

Influence of voriconazole and fluconazole on reconstituted multilayered oesophageal epithelium infected by Candida albicans. J.Bernhardt, H.Bernhardt, M.Knoke, K.Ludwig. Mycoses, 47, 7, p.330, October 2004.

Evaluation of Human Esophageal Epithelium Permeability in Presence of Different Formulations Containing Hyaluronic Acid and Chondroitin Sulphate. Gaia Pellegatta, Marco Spadaccini, Laura Lamonaca, Vincenzo Craviotto, Ferdinando D'Amico, Laura Ceriotti, Marisa Meloni, Alessandro Repici. Medical Devices: Evidence and Research 2020.

In Vitro Modelling of Barrier Impairment Associated with Gastro-Oesophageal Reflux Disease (GERD). Marisa Meloni, Paolo Buratti, Francesco Carriero, Laura Ceriotti. Clinical and Experimental Gastroenterology 2021.

Protective Mechanisms of Liquid Formulations for Gastro-Oesophageal Reflux Disease in a Human Reconstructed Oesophageal Epithelium Model. Laura Ceriotti, Paolo Buratti, Enrico Stefano Corazziari, Marisa Meloni. 2022.

-IVE SkinEthic

DESCRIPTION

Human

NAME

The SkinEthic[™] HVE model is composed of A431 cells (derived from a vulval epidermoid carcinoma) cultivated on an inert polycarbonate filter at the air liquid interface in a chemically defined medium.

This model is histologically similar to the in vivo vaginal mucosa.

Human **Vaginal Epithelium**

FORMAT

0.5 cm²

HTS 24-well plate 0.33 cm²



APPLICATIONS

Vaginal Irritation



Bacterial Adhesion

Omics

SPECIFIC MARKERS

Differentiation markers:

- Keratin
- Involucrin

Vaginal Epithelium

SkinEthic[™] HVE / Human Vaginal Epithelium





REFERENCES

Integrated in vitro vaginal safety approach for bath and body wash products utilizing SkinEthic Human Vaginal Epithelium (HVE) model.

Vinayak S, Alonso A, Bertino B, Costin GE, de Brugerolle de Fraissinette A, Orak D, Inglis H, Kazmi P, Raabe H, Re T. SOT.

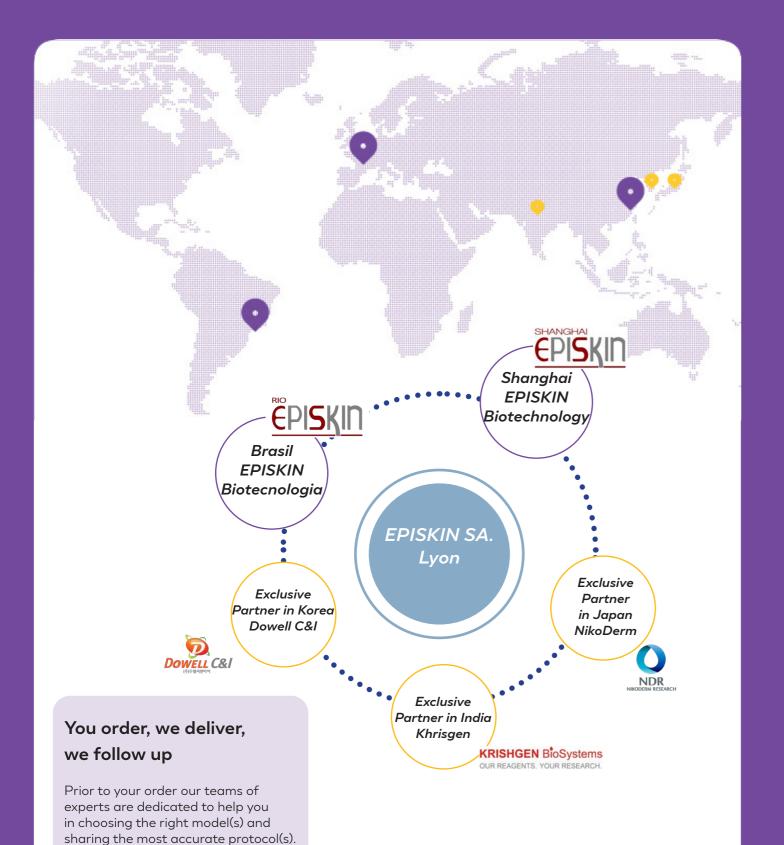
Quantitative expression of the Candida albicans secreted aspartyl proteinase gene family in human oral and vaginal candidiasis. Naglik JR et al. Microbiology, 154, 3266-3280.

Candida albicans-secreted aspartic proteinases (Sap) modify the epithelial cytokine response in an in vitro model of vaginal candidiasis. Schaller M, Korting HC, Borelli C, Hamm G, Hube B.

Infect Immun 73 (5): 2758-2765.

The secreted aspartyl proteinases sap1 and sap2 cause tissue damage in an in vitro model using vaginal candidiasis using reconstituted human vaginal epithelium. Schaller M, Bein M, Korting HC, Baur S, Hamm G, Monod M, Beinhauer S and Hube R

Infection and Immunology, 71, 6, 3227-3234.



Contact us

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ACADEMY

We support our customers

We insure to users of 3D models the best support for implementing 3D's based standard protocols and assist them, for developing new applications of human reconstructed epithelium.

We raise awareness

We relay and illustrate EPISKIN and group's commitments to 3R. Wetrainscientists, KOL, students and future stakeholders to the scientific and regulatory challenges of alternative to animal testing.

Education

EPISKIN Academy created in 2011 is an international program dedicated to promotion and education to alternatives to animal testing methods in toxicology.

As a world leader in tissue engineering, we are committed to share our knowledge and expertise with scientific community to support implementation reconstructed tissues. It is with this in mind that we offer training on validated in these fields.

regulators for the challenges of 21st century toxicology. Certified Program WORLWIDE CERTIFICATION PROGRAMME with more than 200 scientists

We take care of the logistics for an on time delivery by selecting the

more efficient carrier and validating

Upon reception and further we are

reachable real time for any question.

the route when necessary.

ANYTIME, ANYWHERE



We promote scientific expertise

We promote scientific expertise and uses of 3D models by interacting and networking with scientific community. We participate actively to congresses, workshops for sharing scientific results onto human 3D models and promote publication of new data generated by our worldwide





EPISKIN: your reliable and trustworthy partner for your benefits