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# Inflammatory and non-inflammatory mediated itch/pruritus: New Model!





"We combine
our unique expertise,
our project design creativity,
and our passion to advance
our clients' success in
delivering novel and gamechanging skin and hair
research solutions"

Founder & CEO: Prof. Dr. Ralf Paus

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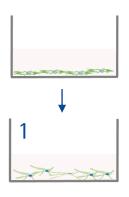
#### For enquiries, please contact:

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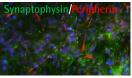
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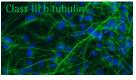
# NEW MODEL: Sensory Reinnervation of Human Skin by Human Neural Stem Cell—Derived Peripheral Neurons *Ex Vivo*

#### Our workflow:



1. Differentiation of human iPSC derived neural stem cells *in vitro* 





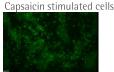
Human iPSC derived neural stem cells start to express: class III β-tubulin which is associated with neuronal maturation and is a microtubule element of the tubulin family found almost exclusively in neurons and neurite extensions (Sainath and Gallo, Cell Tissue Res 2015), peripherin which is a peripheral nervous system neuronal marker (Yuan et al., J Neurosci 2012), and synaptophysin which is a marker of mature neurons (Kwon et al., Neuron. 2011)

Animal-free model!

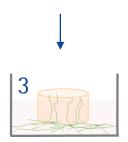
A NEW pre-clinical assay for testing the effects of drugs on innervated skin ex vivo



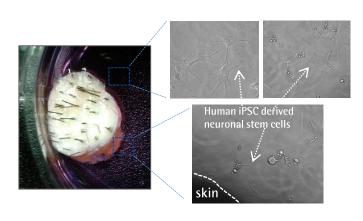




2. Initiation co-culture of human skin with human differentiated iPSC derived neural stem cells *ex vivo* 

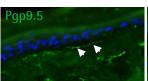




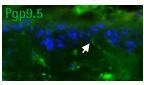


Topical application possible

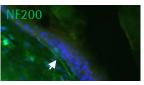
3. Model ready to use: Human skin punch is fully re-innervated



nerve fibers (PGP9.5+) reaching the epidermis



nerve fibers (Pgp9.5+) entering into the epidermis

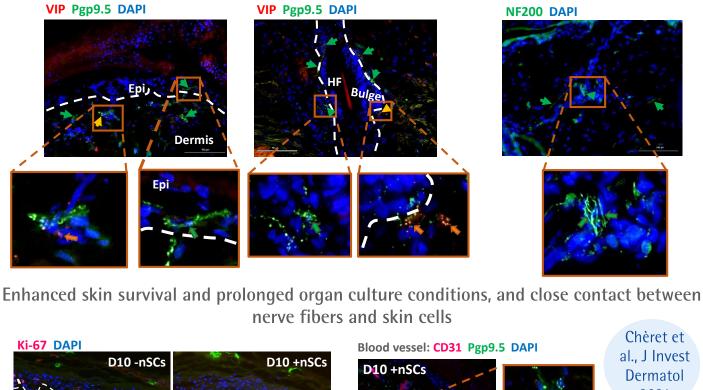


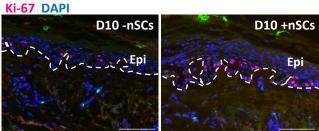
myelinated (NF200+) nerve fibers along the hair follicles

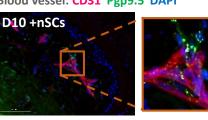
INSTEAD: No remaining nerve fibers can be detected in human skin cultured ex vivo in the absence of human iPSC derived neural stem cells

## **NEW MODEL: Sensory Reinnervation of Human** Skin by Human Neural Stem Cell-Derived Peripheral Neurons Ex Vivo

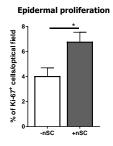
Presence of peptidergic and myelinated sensory fibers after re-innervation

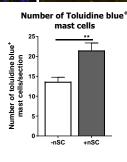


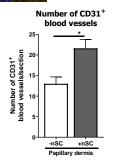


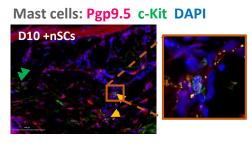


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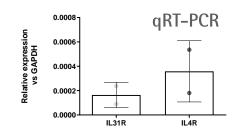




This novel assay can be combined with other diseases models ex vivo (e.g. stimulation with IL-4 and IL-13 for mimicking atopic dermatitis) and can be utilized for dissecting and manipulating the bi-directional communication between defined skin and hair follicle cell populations and (sensory) human nerve fibers under stringently controlled ex vivo conditions, or for testing drugs that target the cross-talk between human skin and hair follicles and cutaneous nerve fibers. Relevant for e.g. sensitive skin, itch, atopic

dermatitis, psoriasis

Differentiated human iPSC-derived Neural Stem Cells (nSCs) also express IL4R and IL31R, and other itchrelevant markers



### WHY US?

Our vision is
to provide our clients and
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from basic science to
translational applied and contract
research of high relevance for
clinical applications.

Great network of dermatologists and plastic surgeons collecting samples from healthy and diseased skin



World-class scientific leadership & international team Clinicallyrelevant ex vivo and in vivo models

Strong
academic
background &
publication
record

#### What we can do for our clients:

- Conceptualize & build proof-of-concept studies
- Carry out full service portfolio for pre-clinical skin & hair research (in vitro, and ex vivo assays, and humanized mouse models)
- Provide access to human healthy & diseased skin and hair specimen
- Develop novel cutting edge methodologies and techniques
- Develop tailor-made & customized assays
- Identify, characterize, or validate novel targets and therapeutics for skin & hair disorders
- Discover mechanistic action stories, biomarkers & predictors of response
- Investigate side effects in the skin or hair follicle
- Conduct investigator initiated skin & hair clinical trials
- Prepare comprehensive project reports & manuscript drafts

Innovation is our passion: Innovative Technology Program

Exceptional state-of-the art research technology

Investigative dermatology:

Acne Vulgaris, Atopic Dermatitis, Psoriasis, Alopecia Areata, Androgenic Alopecia, Hidradenitis Suppurativa, Vitiligo, Chronic Itch, Prurigo Nodularis, etc. We are supported by world-wide recognized experts in dermatology: Alfredo Rossi, Amos Gilhar, Désmond J. Tobin, Erwin Tschachler, Falk G. Bechara, Francisco Jimenez, Kristian Reich, Mauro Picardo, Thomas Luger, Tiago R. Matos, and many more!

Global client list & testimonials

Biobank:
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& hair samples
(patients &
healthy subjects)