NEW METHODOLOGICAL APPROACH TO FOLLOW THE **RE-EPITHELIALIZATION PHASE IN THE WOUND-HEALING PROCESS ON A 3D FULL THICKNESS SKIN MODEL**



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Fallou Bénédicte¹, Pelleter Mylène¹, Innamorato Florence¹, Roche Mickaël¹, Bataillon Michel¹ 1 Episkin, Lyon, France

INTRODUCTION

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RESEARCH & INNOVATION

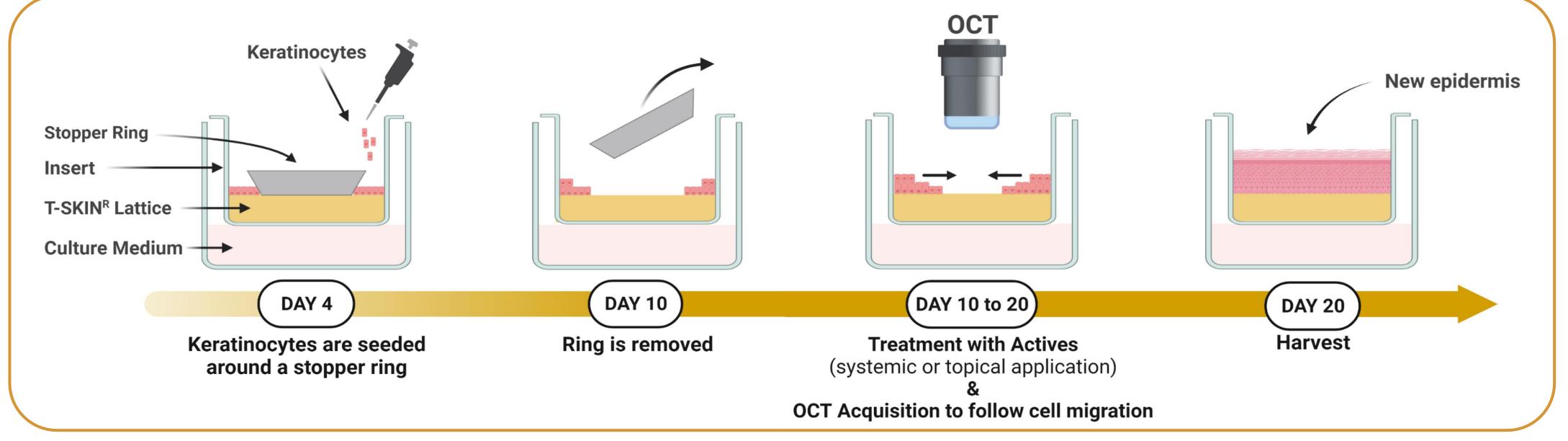
In the *in vitro* evaluation architecture of skin regenerative actives, the different mechanisms of action of ingredients need to be investigated to identify new candidates in skin regeneration, in the context of healing or anti-aging axis. A 3D migration model has been developed to mimic the re-epithelialization phase III and IV of the wound healing process [1]. Thanks to this model, in 2017, a method was developed to analyze the regenerative process using an invasive technology, i.e., Papanicolaou staining and histological observation at each day of the study.



The aim of this new methodological approach was to develop a non-invasive method allowing to follow the reconstruction kinetics of tissues over the different healing steps with a final quantification to be more predictive and consolidate the in vitro evaluation of active ingredients presenting a regenerative potential.

In this migration study, EPISKIN developed a new optical acquisition and processing software based on a non-invasive Optical Coherence Tomography (OCT) approach combined with a final quantification of the histological quality.

MATERIALS & METHODS



DESCRIPTION OF THE MIGRATION TEST

OCT is a non-invasive imaging technique, based on the analysis of infrared light reflected by tissues and on the creation of an interference signal [2]. This technique allows to visualize the surface aspects, the 2D and the 3D structures of the migration model during re-epithelialization and thus to follow the kinetics of closure.

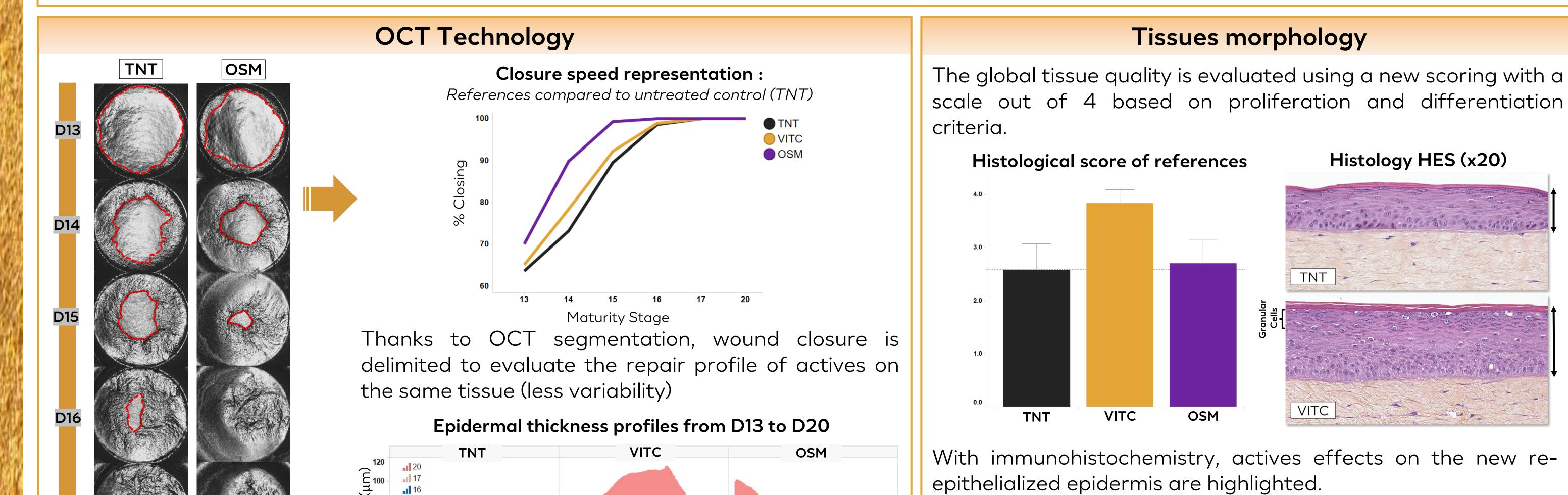
Through acquisition and segmentation with specifics algorithms, epidermal and dermal thicknesses, and epidermis profile of the tissue after reconstruction are also determined.

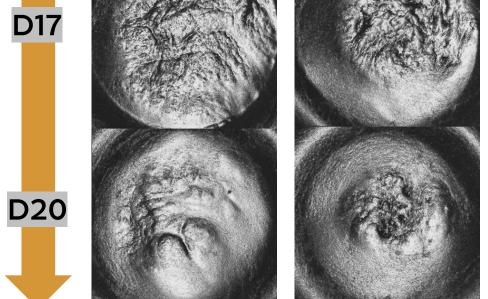
Then, global morphology of the treated tissues is scored at the end of treatment by histology [3].

RESULTS AND DISCUSSION

2 clinical references are used for their positive effect on the test :

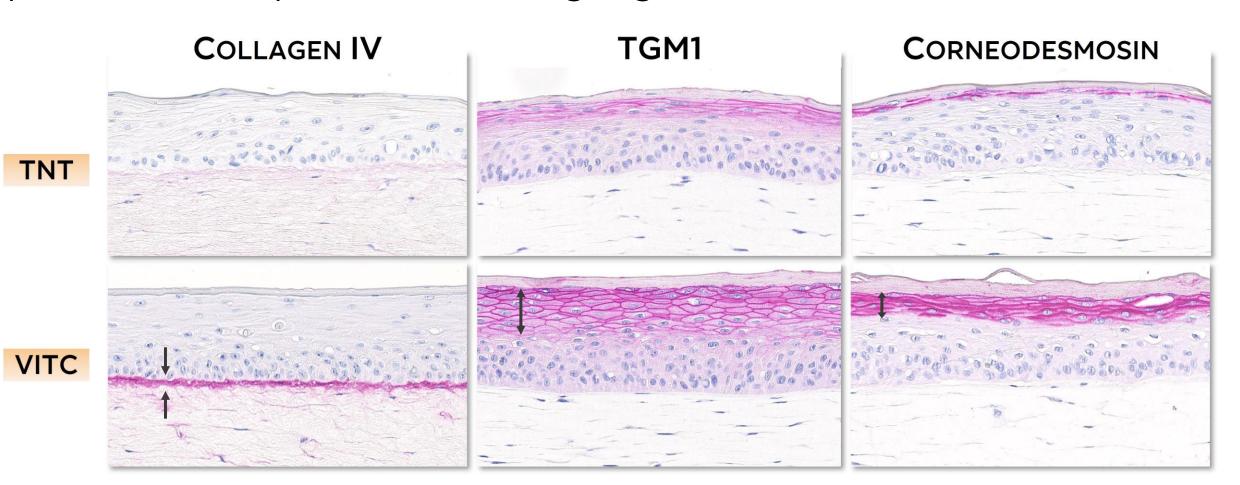
- ✓ Oncostatine M (OSM) increases the speed of migration and proliferation of keratinocytes (eq. wound closure)
- ✓ Vitamin C (VITC) improves quality of reconstruction, organization and differentiation (eq. quality of skin)





Contrast images of the 3D Migration model during re-epithelialization (Untreated TNT vs OSM positive control)

Ring edge Center of tissue Re-epithelialization effect is observed through keratinocytes migration from edge to center of wound (blue profiles) while differentiation step is followed with increasing of epidermal thickness (red profiles).



CONCLUSION 5

Thanks to this work, a new reliable, robust and relevant non-invasive evaluation method has been developed, essential to understand the mechanisms of actions of our actives in the process of skin regeneration and healing.

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