

High relative abundance of bacillales is associated with Epidermolysis Bullosa (EB) at different stages of wound healing

D. L. Balacco¹, A. Bardhan^{1,2}, M. Grant¹, S. A. Kuehne¹, J. Hirschfeld¹, A. Heagerty², I. Chapple¹

¹ Periodontal & Dermatology Research Groups, Institute of Clinical Sciences, University of Birmingham, Birmingham, UK

² Epidermolysis Bullosa Unit, Department of Dermatology, University Hospitals Birmingham National Health Service Foundation Trust, Birmingham, UK

What is Epidermolysis Bullosa (EB)?

EB
Epidermolysis Bullosa is a group of **rare** genetic skin disorders

Incidence of 19.6 per million live births

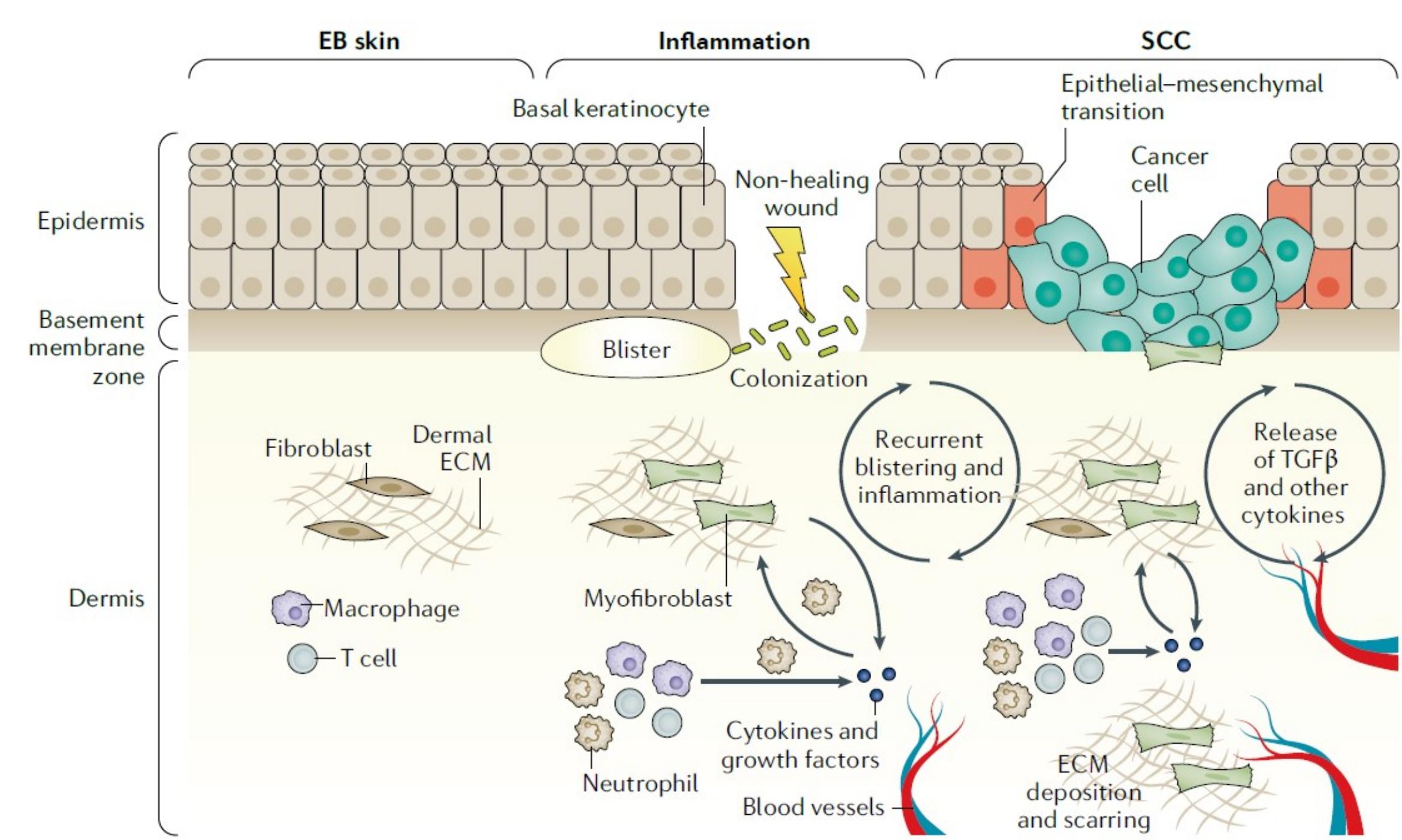
Mucocutaneous fragility and blister formation

Affects internal organs

At least 16 genes involved

There is no cure

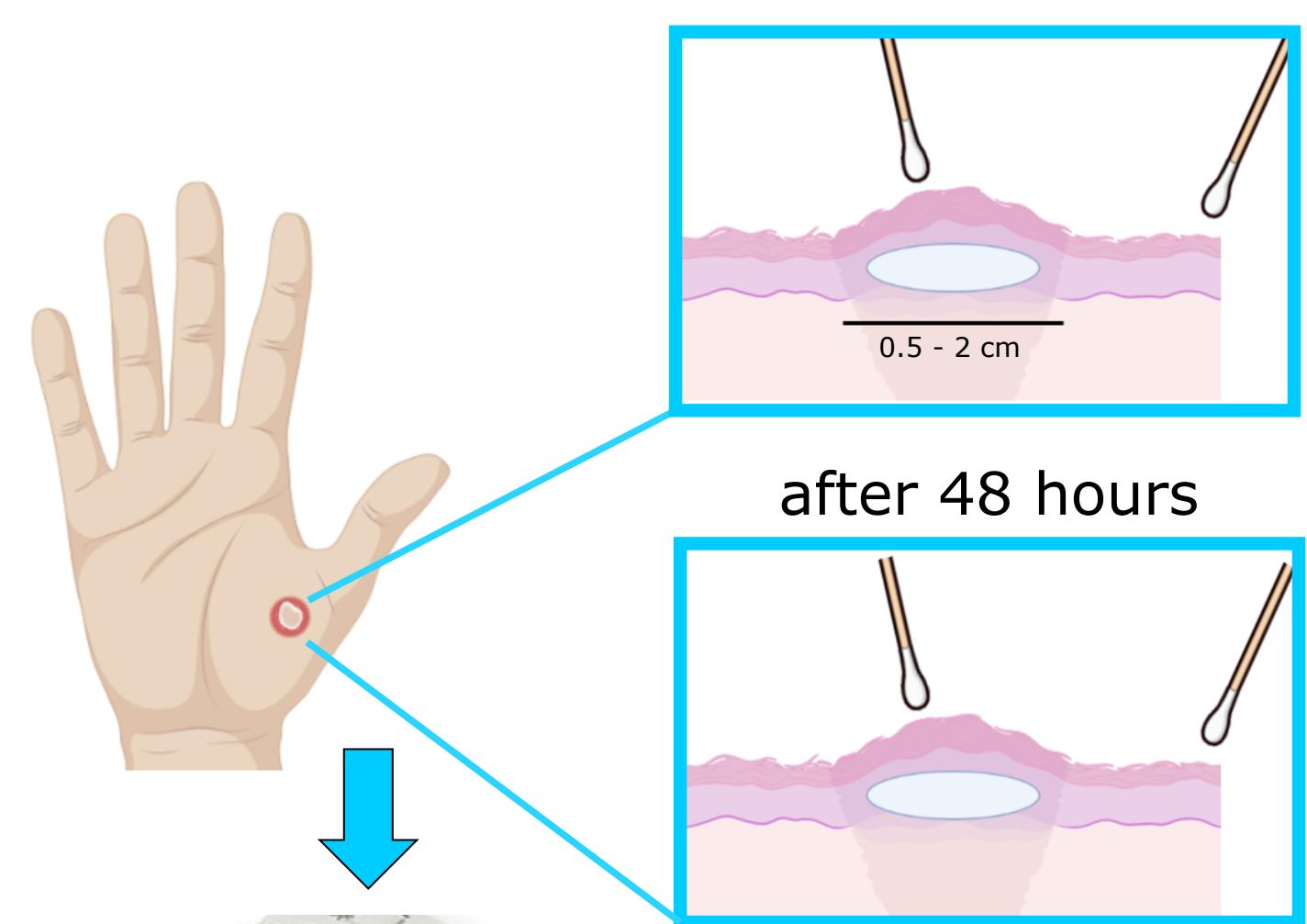
EB is classically grouped into four major types according to the level of skin cleavage:
EB simplex (**EBS**), junctional EB (**JEB**), dystrophic EB (**DEB**), and Kindler EB (**KEB**).
A heterogeneous phenotypic spectrum exists with: persistent blistering, inflammation, scarring, delayed re-epithelialisation, abnormal wound healing, and infection.
It can cause disability and death. Understanding the **skin microbiome** and its relationship to the **inflammatory-immune response** are fundamental prerequisites for developing novel therapies.



Certain types of EB are associated with an increased risk of developing cutaneous squamous cell carcinoma (SCC)¹.

The study aims to characterise, for the first time, the skin microbiome of blisters and unaffected skin of EB patients employing a metagenomic approach.

Methods



- Blister between 0.5 and 2 cm
- 30 seconds skin swab
- Swabs from blistered skin
- Swabs from healthy skin surrounding the blister

MG-RAST
metagenomics analysis server

2

- 1) Bardhan A, et al, Epidermolysis bullosa. Nat Rev Dis Primers. 2020 Sep 24;6(1):78.
- 2) Meyer F, et al., The metagenomics RAST server – a public resource for the automatic phylogenetic and functional analysis of metagenomes. BMC Bioinformatics. 2008 Sep. 19; 9:386.

Figures created using Biorender

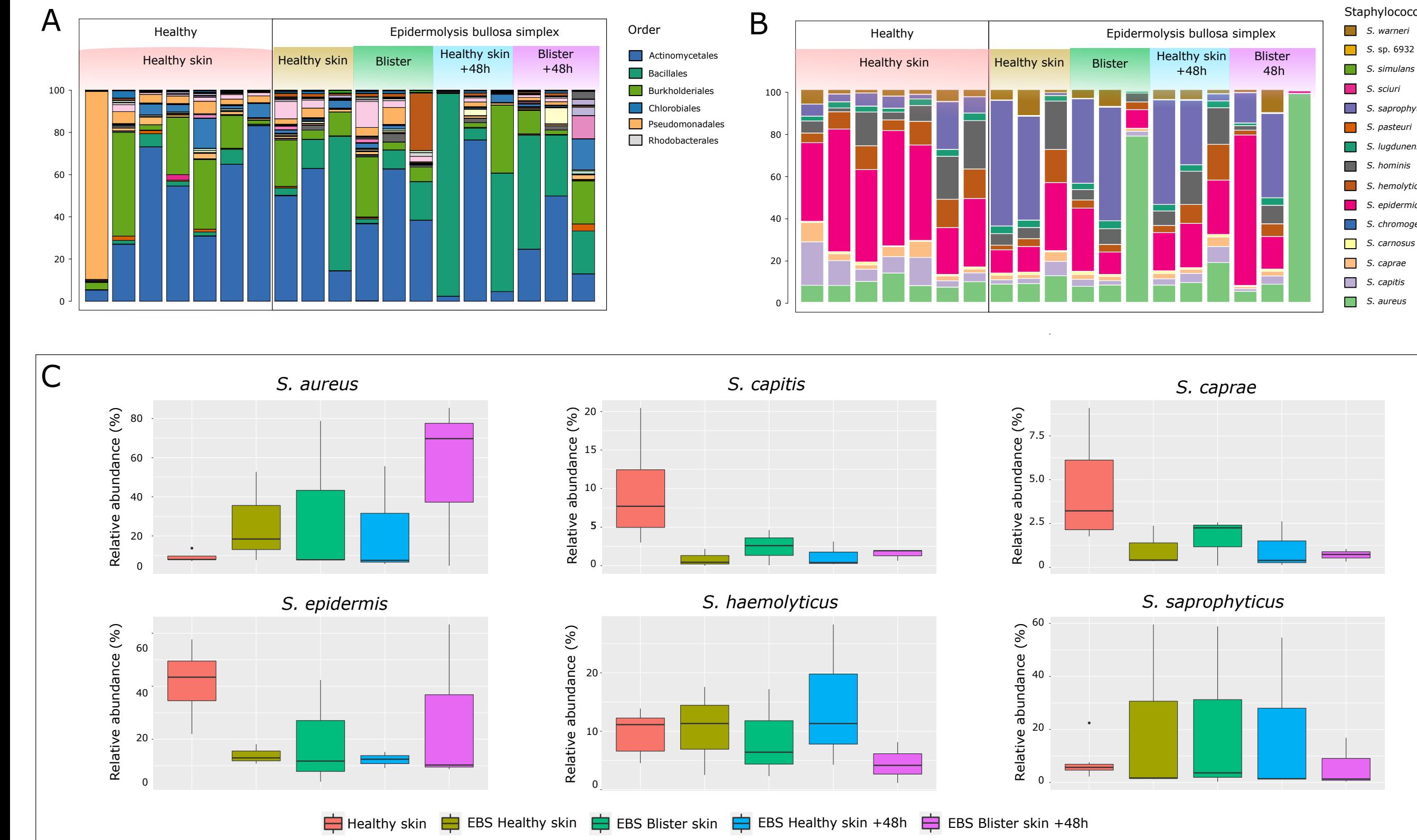
Contacts

d.l.balacco@bham.ac.uk @dariobalacco

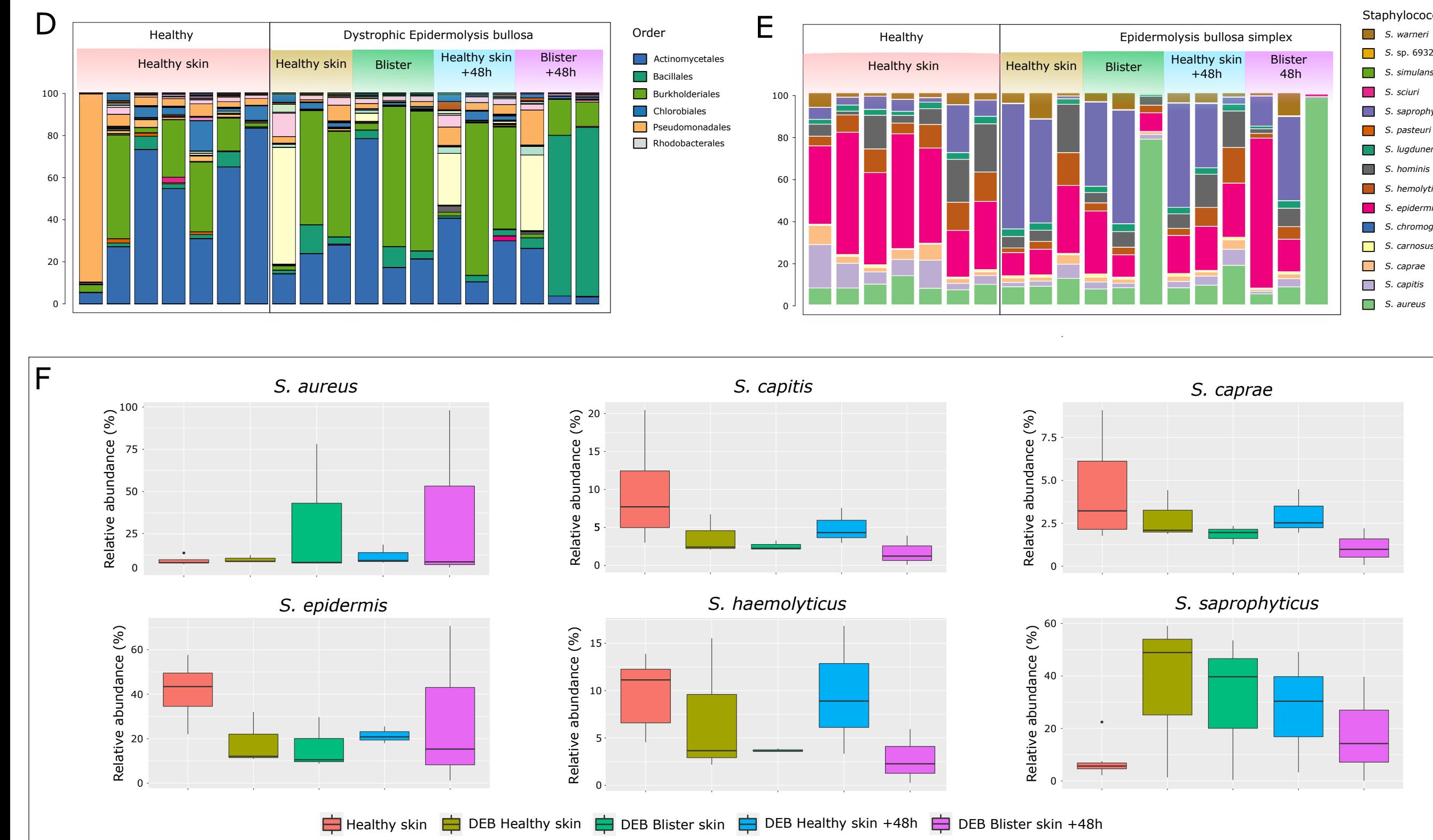
References

Results

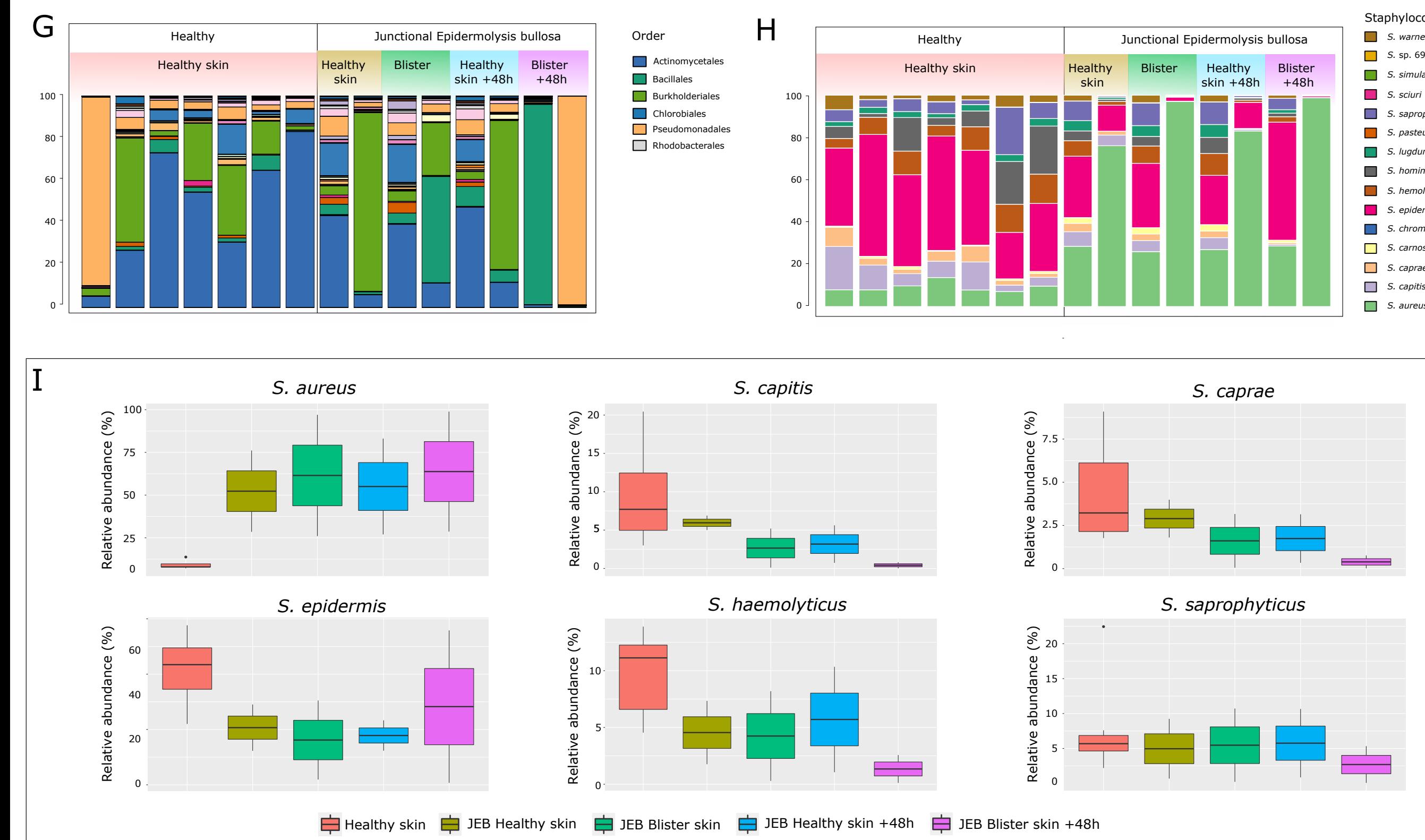
Epidermolysis Bullosa simplex



Dystrophic Epidermolysis Bullosa



Junctional Epidermolysis Bullosa



Bacterial relative abundance in healthy individuals (n=7) and people affected by EB simplex (n=3) (A,B,C), Dystrophic EB (n=3) (D,E,F), and Junctional EB (n=2) (G,H,I). Individual content of bacteria; most abundant bacteria are shown in the legend with order as taxonomical level. B,E,H) Relative abundance of *Staphylococci* species. C,F,I) Relative abundance of the most abundant species of *Staphylococci*.

Conclusion

- The skin microbiome of EB patients has a **high** relative abundance of **Bacillales**.
- Composition of the skin microbiome in EB patients varies **during wound healing**; in particular, the *Staphylococci* showed species-specific and EB type-specific abundancies.
- We observed a **decrease** of *S. epidermidis* in all EB samples.
- In **EBS**, *S. aureus* showed an **increase** of the relative abundance of blistered skin after 48 hours of blister formation.
- In **DEB**, we observed a **high** relative abundance of *S. saprophyticus*, with progressive decrease over time.
- In **JEB**, *S. aureus* maintained a **high** relative abundance in EB patients skin samples.
- These results suggest that there is **association** between shifts in the microbiome and **wound healing** in EB patients subtypes, which can be **exploited therapeutically**.