The 1st International Online Conference on Fermentation

MDPI

12-13 November 2025 | Online

Hacking gut-immune axis with non-Saccharomyces yeasts: effects of Candida intermedia supplementation in SARS-CoV-2-vaccinated animals

Renan E. A. Piraine^{1,2}, Neida L. Conrad¹, Vitória S. Gonçalves¹, Jeferson V. Ramos¹, Júlia L. Froldi², Fausto Almeida², Fábio P. L. Leite¹

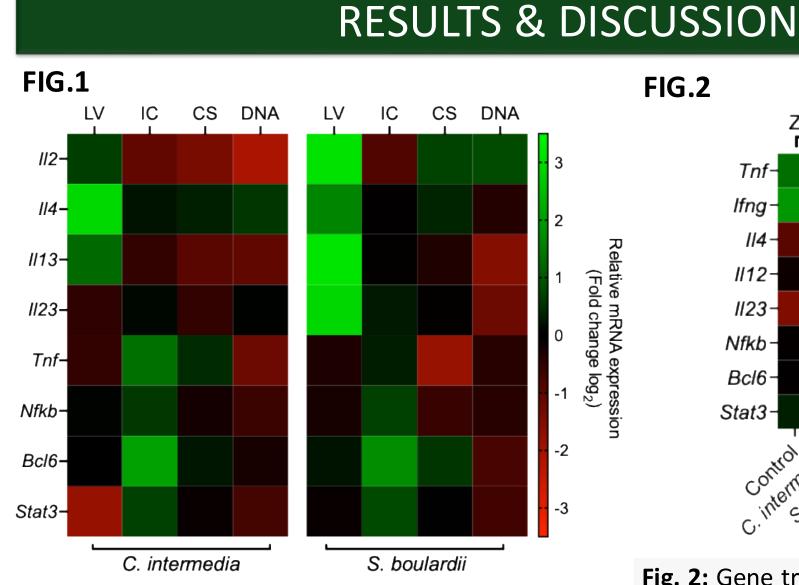
1 Microbiology laboratory, Technological Development Center, Federal University of Pelotas, Pelotas, Brazil
2 Department of Biochemistry and Immunology, Ribeirão Preto Medical School, University of São Paulo, Ribeirão Preto, São Paulo, Brazil

INTRODUCTION & AIM

Non-Saccharomyces yeasts are emerging as promising probiotics, showing biological effects comparable to the reference strain *Saccharomyces boulardii*. In previous studies, we identified *in vitro Candida intermedia* ORQ001 (a non-albicans yeast not considered a common human pathogen) as possessing probiotic traits [1]. Probiotics are known to modulate host immune responses by enhancing antibody production, increasing phagocytic activity, and altering cytokine expression. Based on these findings, this study aimed to evaluate the immunomodulatory effects of *C. intermedia* in mice vaccinated with inactivated SARS-CoV-2, providing further evidence of its probiotic potential.

METHOD Procedure 35 Days 28 -5 -4 -3 -2 -1 16 17 18 19 20 O Blood and fecal samples •••• Supplementation Euthanasia of animals Vaccination S. boulardii CNCM-1745 C. intermedia ORQ001 Immune system cells culture **ELISA** Macrophages (RAW 264.7) IgG, sIgA Splenocytes (ex vivo)

Transcription factors Nfkb, Bcl6, Stat3

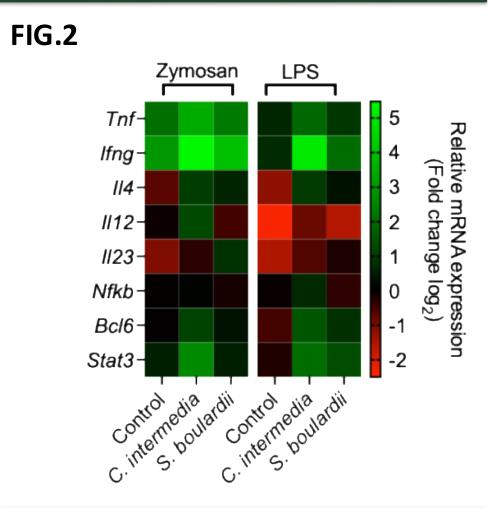


Real-time PCR

Il2, Il4, Il12, Il13, Il23, Tnf, Ifng

Cytokines

FIG. 1: Gene transcription of cytokines (*II2, II4, II13, II23, Tnf*) and transcription factors (*Nfκb, Bcl6, Stat3*) in RAW 264.7 macrophages stimulated with yeasts and their derivatives. LV: live cells; IC: inactivated cells; CS: culture supernatant; DNA: DNA extracted from fungal cultures.

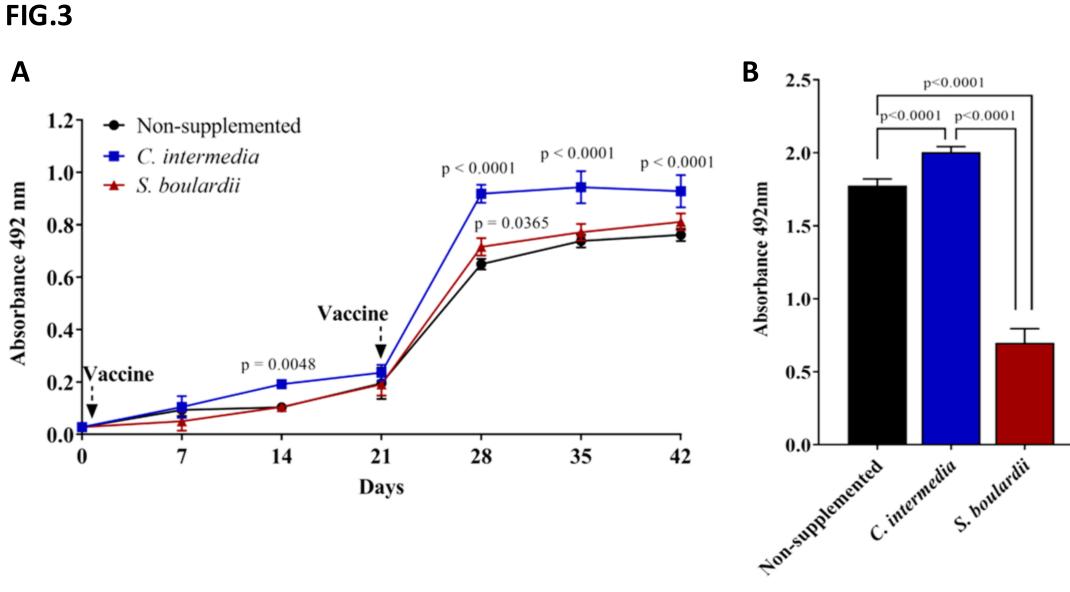


Microbiome

Bacteria (16S)

METAGENOMICS

Fig. 2: Gene transcription of cytokines and transcription factors during splenocyte stimulation. Splenocytes were obtained from animals supplemented with *C. intermedia* or *S. boulardii*, or from nonsupplemented). Cells were cultured in vitro and stimulated with Zymosan or LPS.



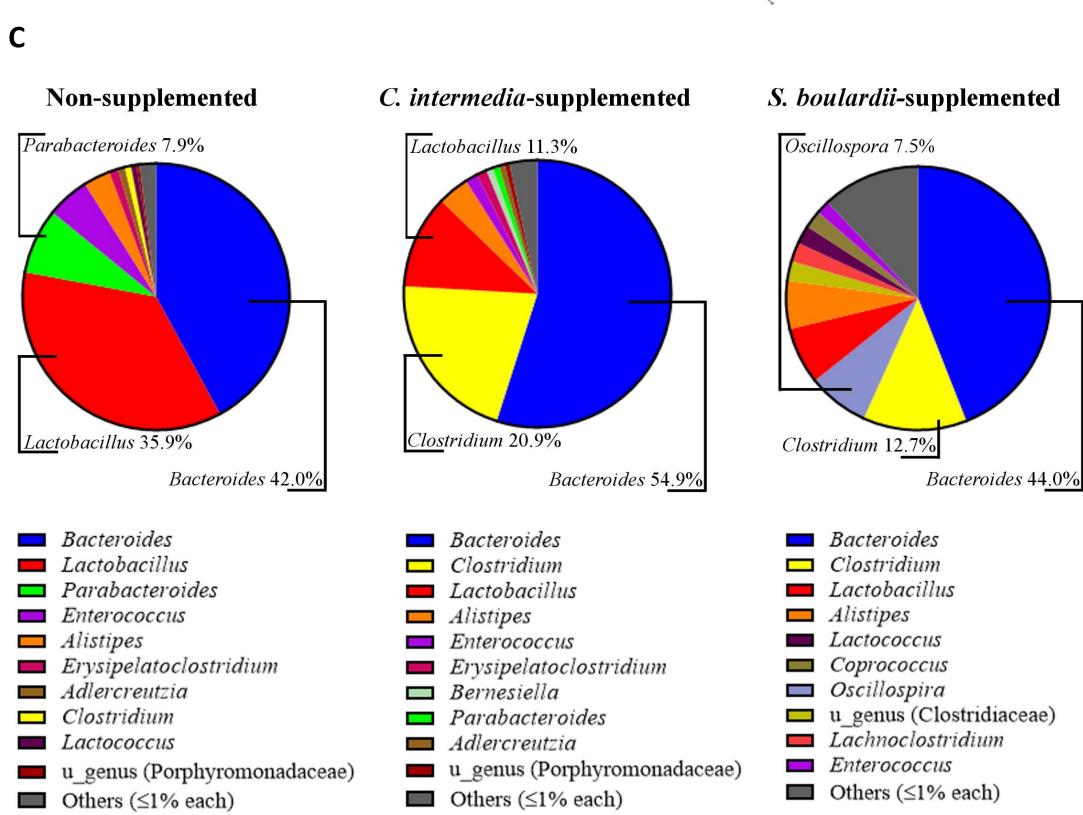


FIG. 3: Humoral immune response and gastrointestinal tract microbiome of non-supplemented animals (control) and supplemented with *C. intermedia* and *S. boulardii*, vaccinated with inactivated SARS-CoV-2. A) Dynamic of serum IgG production, analyzed each seven days from pooled sera of the experimental groups. B) sIgA detection on pooled fecal samples of mice in the 42nd experimental day. C)The most abundant bacterial genera in the sequenced samples, emphasizing the top 3 in each sample.

CONCLUSION

C. intermedia supplementation modulated the expression of key immune-related genes and enhanced humoral responses in mice. Furthermore, its influence on gastrointestinal microbiota suggests a synergistic effect on vaccine immunogenicity. This work provides original insights that may broaden the understanding of host—microbe interactions in vaccination and contribute to the development of innovative probiotic-based strategies to improve vaccine efficacy.

FUTURE WORK / REFERENCES

[1] Piraine, R.E.A.; Retzlaf, G.M.; Gonçalves, V.S.; Cunha, R.C.; Conrad, N.L.; Bochman, M.L.; Leite, F.P.L. Brewing and probiotic potential activity of wild yeasts *Hanseniaspora uvarum* PIT001, *Pichia kluyveri* LAR001 and *Candida intermedia* ORQ001. *Eur. Food Res. Technol.* **2023**, *249*, 133–148.

ACKNOWLEDGMENTS

We thank Conselho Nacional de Desenvolvimento Científico e Tecnológico (CNPq) for the financial support, process number 50538/2021-9