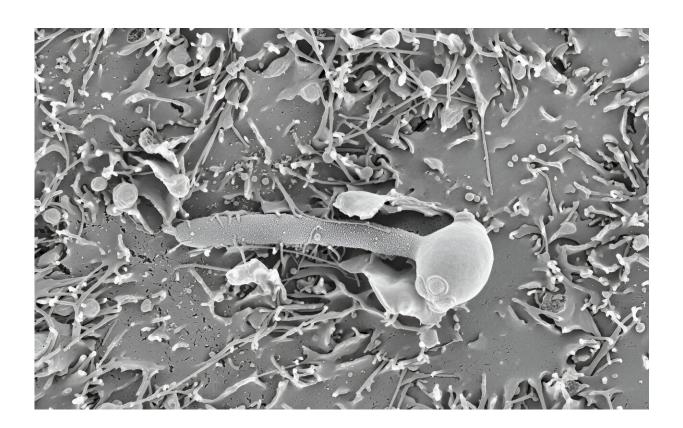


Candida evolution disclosed: New insights into fungal infections

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A Candida albicans cell forming a hyphal extension. Credit: IRB Barcelona

Global fungal infections, which affect 1 billion people and cause 1.5 million deaths each year, are on the rise due to the increasing number of medical treatments that heighten vulnerability. Patients undergoing chemotherapy or immunosuppressive treatments after organ transplant



often present compromised immune systems.

Given the emergence of resistant strains, the limited variety of current <u>antifungal drugs</u> as well as their cost and side effects, the <u>treatment</u> of these infections is challenging and brings about an urgent need for more effective treatments.

In this context, a team from the Institute for Research in Biomedicine (IRB Barcelona) and the Barcelona Supercomputing Center—Centro Nacional de Supercomputación (BSC-CNS), led by the ICREA researcher Dr. Toni Gabaldón, has identified hundreds of genes subject to recent, clinically-relevant selection in six species of the fungal pathogen, Candida. The findings are <u>published</u> in the journal *Nature Microbiology*.

"This work highlights how these pathogens adapted to humans and antifungal drugs and provides valuable knowledge that could lead to better treatments for Candida infections," explains Dr. Gabaldón, head of the Comparative Genomics lab at IRB Barcelona and the BSC.

More than 2,000 genomes from six different species

The study delves into the evolutionary landscape of Candida pathogens by analyzing approximately 2,000 genomes from clinical samples of six major Candida species. These genomes are stored in public databases. The researchers compared these genomes to a reference, creating a comprehensive catalog of genetic variants.

Building on previous work addressing drug-resistant strains, the researchers conducted a Genome-Wide Association Study (GWAS) to identify genetic variants linked to antifungal drug resistance in clinical isolates. This approach provided insights into both known and novel mechanisms of resistance toward seven antifungal drugs in three



Candida species.

"Additionally, a concerning finding has arisen from the study: the potential spread of resistance through mating between susceptible and resistant strains, contributing to the prevalence of drug-resistant Candida pathogens," explains Dr. Miquel Àngel Schikora-Tamarit, a postdoctoral researcher in the same lab and first author of the study.

In addition, by focusing on variants acquired recently among clinical strains, the researchers detected shared and species-specific genetic signatures of recent selection that inform on which adaptations might be needed to thrive and spread in human-related environments.

Beyond the novel insights into the adaptation of Candida, the study provides a valuable resource, namely a comprehensive catalog of variants, selection signatures, and drivers of drug resistance.

This knowledge not only contributes to our understanding of these infections but also lays the groundwork for future experiments and potential advancements in the development of more effective treatments for Candida infections.

More information: Miquel Àngel Schikora-Tamarit et al, Recent gene selection and drug resistance underscore clinical adaptation across Candida species, *Nature Microbiology* (2024). DOI: 10.1038/s41564-023-01547-z

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