

**TITLE:** CANDIDA PARAPSILOSIS SPECIES COMPLEX SUSCEPTIBILITY TO ANTIFUNGAL AGENTS AND GERANIOL.

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**ABSTRACT:**

*Candida parapsilosis* species complex, composed by *C. parapsilosis sensu stricto*, *C. orthopsilosis* and *C. metapsilosis*, represents a recognized threat in the nosocomial environment by forming drug-recalcitrant biofilms on catheters and other medical devices. In Brazil, the *C. parapsilosis* species complex is responsible for 15-30% of systemic infections. The use of natural components with antifungal activity has been shown to be a promising therapeutic alternative in infections by *Candida* spp. Among these compounds, the monoterpenoid alcohol geraniol highlight. This study aimed to evaluate the susceptibility of the *C. parapsilosis* species complex strains to antifungal agents and geraniol. Therefore, 37 strains of human, veterinary and environmental sources were used from the fungal collection of Center for Medical Mycology (Fortaleza-Ceará, Brazil), being 21 (56.7%) *C. parapsilosis strictu sensu*, 9 (24.3%) *C. metapsilosis* and 7 (19%) *C. orthopsilosis*. The susceptibility assay was conducted by the broth microdilution method, according to the guidelines of document M-27 (CLSI, 2017). Microdilution was performed in 96-well U-bottom plates in RPMI 1640 medium and inoculum standardized at 0.5 of McFarland scale. Amphotericin B, itraconazole, voriconazole were used at 0,0313 – 16 µg/mL, fluconazole at 0.125 – 64 µg/mL, caspofungin at 0.015 – 8 µg/mL, as geraniol at 4 – 1024 µg/mL. The controls were *C. krusei* ATCC 6258 and *C. parapsilosis* ATCC 22019. The minimum inhibitory concentration capable of inhibiting 50% of fungal growth (MIC50) was determined from the visual reading after 24-48 hours for all drugs. The MIC50 concentration range observed for the drugs were: Amphotericin B (0.0313 – 0.5 µg/mL), itraconazole (0.0313 – 0.25 µg/mL), voriconazole (0.0313 – 0.25 µg/mL), fluconazole (0.125 – 4 µg/mL) and caspofungin (0.125 – 1 µg/mL). All strains evaluated were sensitive to geraniol (64 – 256 µg/mL), with the species *C. metapsilosis* being the most susceptibility, while *C. parapsilosis strictu sensu* presented the highest MIC50 values. Knowledge of the susceptibility to antifungal drugs and alternative compounds provides a better understanding about the different therapeutic possibilities.

**Keywords:** Natural components, antifungal activity, *Candida*.

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