

## The virulence potential of *Prototheca* microalgae explored by in vivo studies in a murine model

Angelika Proskurnicka, Patryk Mazur, Tomasz Jagielski\*

Department of Medical Microbiology, Institute of Microbiology, Faculty of Biology, University of Warsaw, Poland

\* Correspondence to: <u>t.jagielski@uw.edu.pl</u>

unicellular, **Prototheca** are photosynthetic, yeast-like microalgae that inhabit natural variety environments. These microorganisms are the only known plants involved in a number of clinically relevant opportunistic infections vertebrates, collectively known as protothecosis. Very little is known about the pathogenesis of Prototheca spp., including pathogen-host interactions. Thus, the aim of this work was to investigate the virulence **Prototheca** potential the algae by assessing their ability to induce local or systemic infections in an experimental murine model.

## Methods

Type strains of three pathogenic (P. wickerhamii, P. bovis, and P. ciferrii) and one saprophytic (P. stagnora) species were used to experimentally inoculate mice of both immunocompetent and immunodeficient phenotype. The study was carried out on 54 groups of 10-week-old, female mice (6 individuals per each) depending on the inoculum (algae or PBS as a control), the challenging dose (i.e.  $5x10^6$  or  $5x10^7$  CFU/mL), and inoculation route (subcutaneous (S.C.), intramammary (I.M.)), and intraperitoneal (I.P.)). Six weeks post-infection, the animals were euthanized, and their organs were explanted, weighted, and sent for microbiological and histopathological examination.

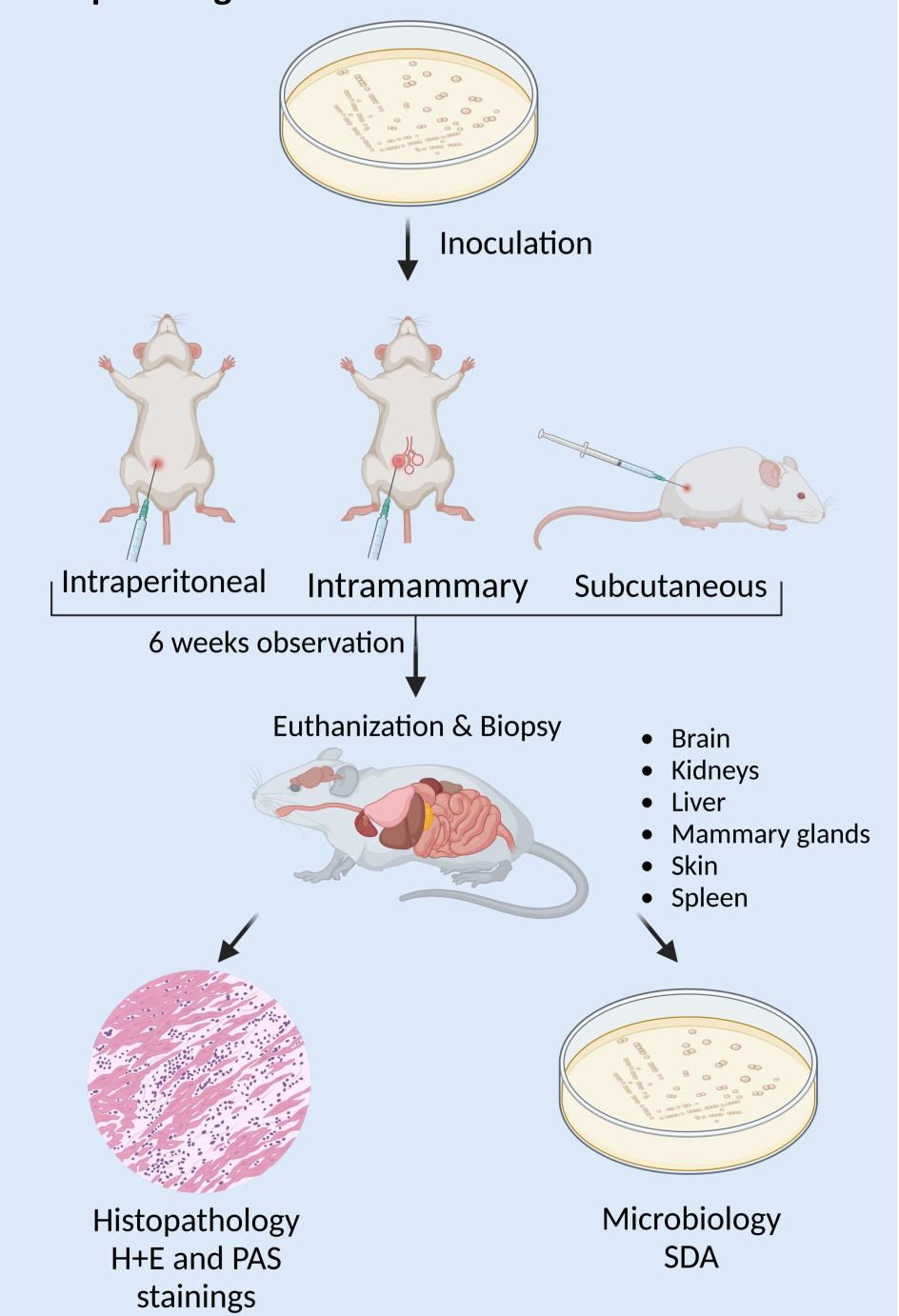
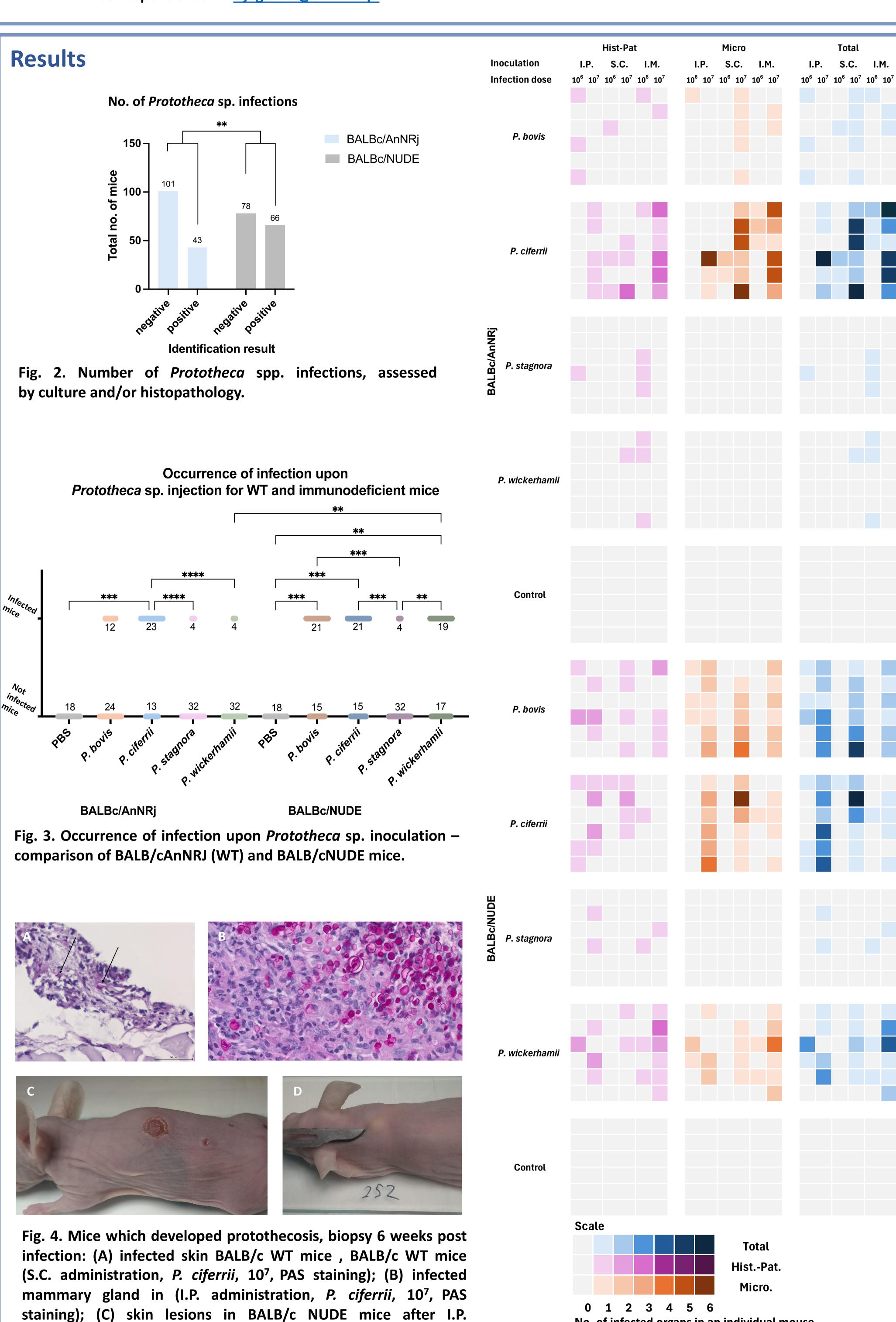


Fig. 1. *Prototheca* spp. infection model.

## Acknowledgement

This research was funded by the National Science Centre within the "OPUS" funding program (contract number: 2019/33/B/NZ6/01283).



the injection site in BALB/c NUDE mice (S.C. administration, Fig. 5. Comparison of histopathology and microbiology for the detection of *Prototheca* sp. infection.

No. of infected organs in an individual mouse

A third (29.9%) of wild-type mice and nearly half (45.8%) of immunodeficient animals showed signs of infection. P. ciferrii accounted for the majority of infections (40.4%), followed by P. bovis (30.3%), and P. wickerhamii (22%). Only 7.3% of infections were due to P. stagnora. All routes of inoculation resulted in a similar number of infected individuals, with the infection efficiency ranging from 31.2% to 35.8%. Mammary glands were the most frequently affected organs, with a prevalence of 53.2% in both healthy and immunodeficient animals. Microbiological culture had a higher detection rate of infection (75.2%) compared to histopathological analysis (66.1%). The most notable inflammatory changes were observed in the spleen and kidneys, while the brain and liver were the least affected.

## Conclusions

*P. bovis*, 10<sup>7</sup>).

☐ The highest virulence potential was observed for *P. ciferrii*.

administration (*P. wickerhamii*, 10<sup>7</sup>); (D) subcutaneous abscess at

- ☐ Immunodeficient mice were more prone to develop *Prototheca* infection.
  - Microbiological culture was more sensitive than histopathology in detection of *Prototheca* sp. infections.