

INVESTIGATION TOWARDS THE SYNTHESIS OF ALKYL C-GALACTOFURANOSIDES DERIVATIVES WITH POTENTIAL ANTILEISHMANIAL PROPERTIES

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Leishmaniasis, a disease caused by protozoan parasites of the genus *Leishmania* and transmitted by the bite of infected sandflies, poses a significant global health challenge. Although this infection is a treatable and curable disease, its treatment faces many challenges. Current drugs suffer from limited availability, side effects, long treatment durations, high costs, and drug resistance issues. Therefore, there is a critical need for the development of new antileishmanials that are not only more effective but also safe and affordable. The abundance of rare carbohydrates assembly with β -D-galactofuranoside linkage within the glycocalyx of these microorganisms [1] led to the exploration of their biochemical pathways as promising antimicrobial targets. Inspired by antimicrobial-active octyl β -D-galactofuranoses [2], hydrolytically resistant alkyl C-galactofuranosides were designed as promising antimicrobial agents. The main synthetic approach to access β -D-galactofuranosyl lipids and other analogs relied on the C-1 allylation of glycosyl acetates, followed by cross-metathesis with various α -olefines and hydrogenation (Figure 1). 22 derivatives were isolated and their biological evaluation against both *Leishmania major* promastigote and *L. mexicana* amastigote revealed antileishmanial activity of 1-(glycosyl)heptadecanes, exhibiting IC_{50} values of $\sim 50 \mu M$ [3]. Notably, 1-(β -D-arabinofuranosyl)heptadecane and its enantiomer showed promising antimicrobial efficacy with IC_{50} values and selectivity index comparable to the conventional antileishmanial drug miltefosine, highlighting their promising therapeutic efficacy.

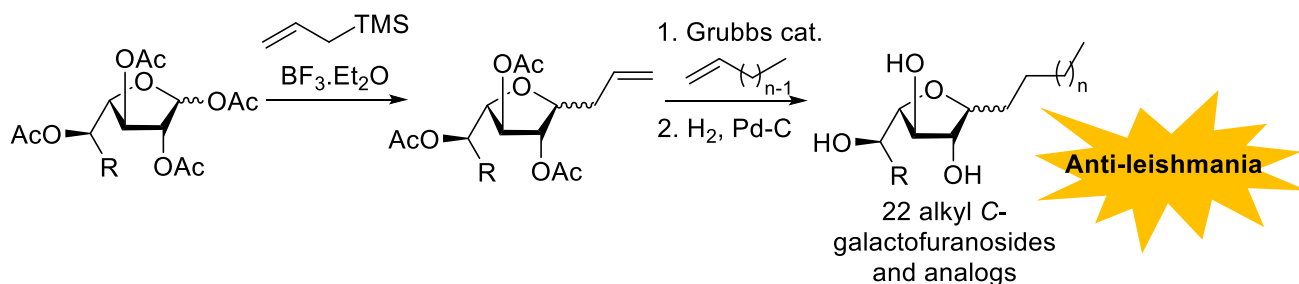


Figure 1

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