

Quinazolinone Amination en route to Compound Library Preparation for the Study of Chagas Disease

Haleigh Rzonca, Dr. Kim Kelly, Huang Yen-Po, Daniel Tolas

Introduction

Chagas disease – Trypanosoma cruzi

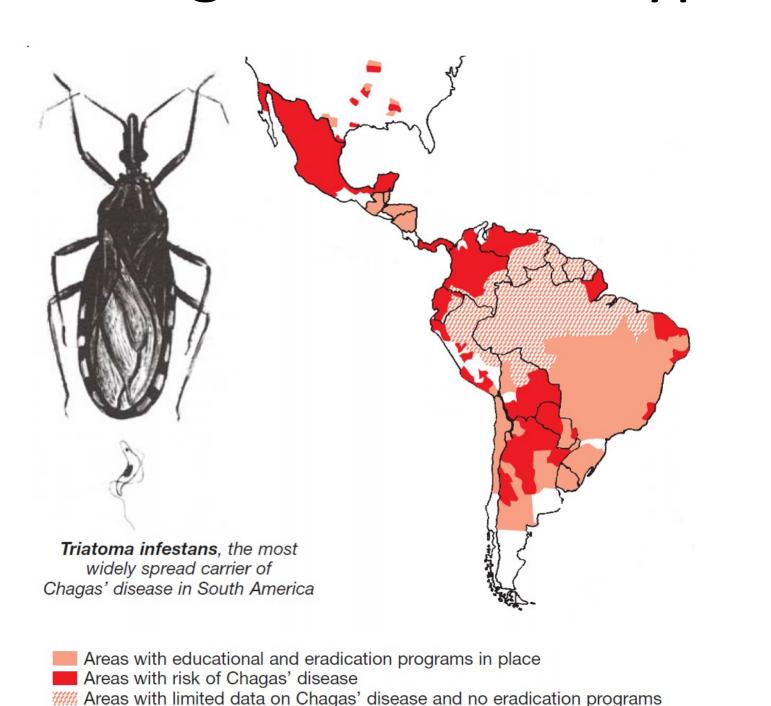
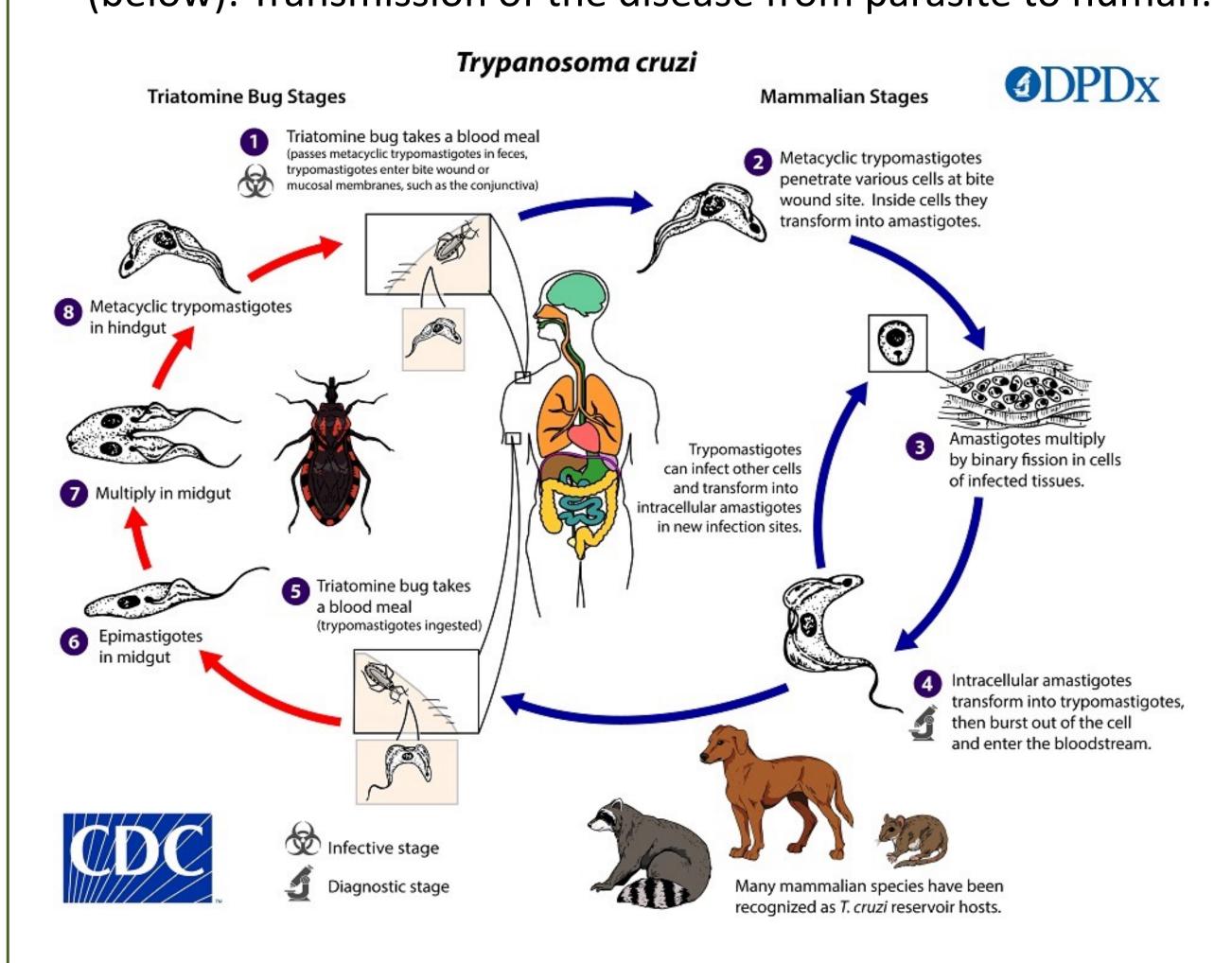


Figure 1: A map of where chagas is present as well as which areas have programs in place. Left is T. cruzi and the parasite that carries the disease.

Map source: PAHO

Currently 8 million people are affected by this disease. There is 2 treatments available, and they only treat acute patients. Acute symptoms include fever and swelling around bitten area. If left untreated the chronic symptoms include a megacolon, megaesophagus, and cardiomyopathy. Figure 2 (below): Transmission of the disease from parasite to human.



METHODS AND MATERIALS

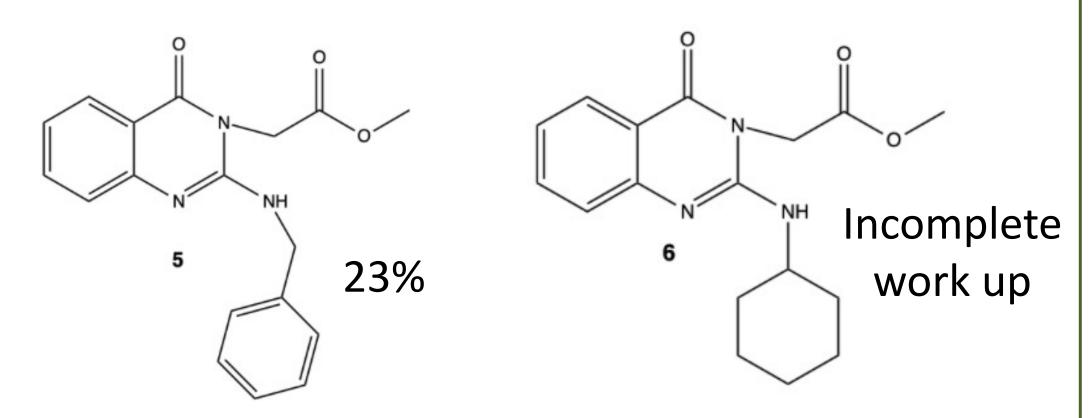
Stage 1: Making quinazolinone

Stage 2: Alkylation
Stage 3: Amination

Amines used: Pyrrolidine, Benzylamine (5), Cyclohexylamine (6)

1: Precursor 2: Quinazolinone 3: Alkyl- Quinazolinone 4: Pyrrolidine
For amination, reagents were placed in a Schlenk
bomb fitted flask, heated to 100 °C and stirred for
approximately 3-6 hours. Thin layer
chromatography were taken after an hour and
then every 30 minutes until completion. H-NMR
data indicated whether the desired product was
created based on H resonance. We experimented
with 3 different amines.

RESULTS



Although we have low yields, our work proved that we can create the expected product.

<u>Future</u>

- Optimize amination process
- Increase compound library
- More 1° vs. 2° amines
- Experiment with cyclic vs. acyclic amines
- Experiment with aromatic vs. aliphatic amines

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REFERENCES

Alonso-Padilla J, Rodr iguez A (2014) High Throughput Screening for Anti—Trypanosoma cruzi Drug Discovery. PLoS Negl Trop Dis 8(12): e3259. doi:10. 1371/journal.pntd.0003259 CDC - dpdx - american trypanosomiasis. Centers for Disease Control and Prevention. 2021 Jun 16 [accessed 2022 May 29]. https://www.cdc.gov/dpdx/trypanosomiasisamerican/index.html Hendsbee D, Westmacott C. Chagas Disease: What is it and why should you care? 2018 Dec 18 [accessed 2022 May 29]. https://www.iamat.org/blog/chagas-disease-what-is-it-and-why-should-you-care/

Rassi A, Rassi A, Marin-Neto JA. Chagas disease. The Lancet. 2010;375(9723):1388–1402. doi:10.1016/s0140-6736(10)60061-x