

# Thesis structure comments

## Contents

- Title: “*Exploring evolution using Saturated Transposon Analysis In Yeast*”
  - Comments:
    - \* The title does not convey the right picture of what you did in the thesis.
    - \* You should focus more on the GI part to understand a particular adaptive evolutionary pathway, like the one from  $\Delta$  bem1 .
    - \* An alternative could be: “*Genome-wise quantification of genetic context influence on gene deletions in budding yeast*” **The case of the bem1-bem3 pair in the context of adaptive evolution.**

### 1. Introduction

1. The  $\Delta$ bem1 adaptive evolutionary trajectory
2. Genetic interactions in yeast
3. State of the art (**of what??**)
  - here you can also put , state of start approaches to measure GI in yeast.
4. SATAY as an alternative technique to compute GI
  - Description of the technique
  - why could it be a good candidate to quantify GI in yeast in a high-throughput manner.
5. Introduction to the research problem
  - Intro of what we still dont know
  - The relevance of tackling that question to gain understanding in how biological networks do change during evolution.
6. Research questions
7. Hypotheses
8. Goals
  - Experimental goals
  - Modelling goals

### 2. Materials & Methods

1. Yeast strain construction
  - Construction of a bem1 bem3 strain
2. SATAY
  1. Library formation
  2. DNA extraction
  3. DNA sequencing
    - DNA Digestion
    - DNA ligation
    - PCR

3. Media
4. OD measurements
  - nanodrop machine
5. Population growth assays
  - Biotek protocols
6. Data analysis
  - Population growth assays
  - SATAY library complexity
  - Access to the software

### 3. Model

1. Current knowledge
  - Interactors of bem1
  - Interactors of bem3
  - Interaction type between bem1 and bem3 , when bem3 is knocked down after a bem1 deletion.
    - Change in fitness from population growth data from  $\Delta$ bem1 genotype to  $\Delta$ bem3 $\Delta$ bem1.
  - Interaction type between bem1 and bem3 , when bem1 is knocked down after a bem3 deletion.
    - Change in fitness from population growth data from  $\Delta$ bem3 genotype to  $\Delta$ bem3 $\Delta$ bem1.
2. What is predicted to change after bem1 deletion in WT and in  $\Delta$ bem3 background.
  - focus on the genetic interactors from bem1 in both backgrounds
3. What is predicted to change after bem3 deletion in WT and in  $\Delta$ bem1 background.
  - focus on the genetic interactors from bem3 in both backgrounds
4. Expanding on previous systems to know which lethal deletions of  $\Delta$ bem3 $\Delta$ bem1 are not lethal in  $\Delta$ bem3
  - From Wessel results using ML on trying to predict gene essentiality using a different set of genes
  - Look at common observables that can be extrapolated to our system , like common go terms, common interactors , or functional enrichment of predicted genes in relation with the gene of interest.

### 4. Results

1. Experimental results
  - Complexity of the satay library
  - Sequencing results
2. Modelling results
  1. Prediction on what is the effect of bem1 deletion on a network that do not have bem3 , based on what is known in WT.
  2. Prediction of which type of genes are more prone to be SL in  $\Delta$ bem1 $\Delta$ bem3 background that are not in the single knockouts backgrounds.

### 5. Conclusion & Discussion

- What do we learn after doing this research?
- Do we accomplish what we proposed to do at the beginning?
- How our findings align with our initial hypotheses?
- Implications of the results for our new understanding on how biological networks do change during evolution.

## 6. Future research

- What is still left and yet relevant to do?

## 7. References

## 8. Appendix

- A Protocols
- B Primer sequences
- C Strains
- D Python code