New data analysis and development of new GRNsight functionality will aid researchers in discovering more about gene regulatory networks

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- Coding new GRNsight functionality and performing yeast data analysis can lead to new insights in gene regulatory research.
- Achieving Individual milestones contributes to group effort.
- Cold shock regulation differs between gene clusters.
- Right click function links to a new gene page which helps biologists find information quickly.
- This team's work will aid future research
- There are many avenues available for further research.

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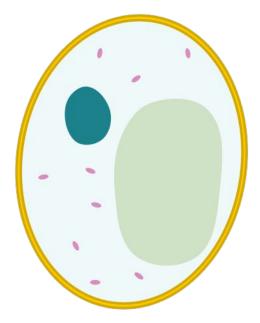
Organisms have mechanisms that allow them to withstand environmental stresses

- Organism -- Shivering
- Organ Systems -- Endocrine, Circulatory
- Organ -- Digestion
- Tissue -- Muscle movement
- Cellular -- Cold shock proteins



Yeast is an ideal model organism for learning about Eukaryotic cells

- Significant growth in a matter of days
- Easy to manipulate in lab
- Can resist and respond to many different environmental stressors
- Controls cell division in a way similar to ours

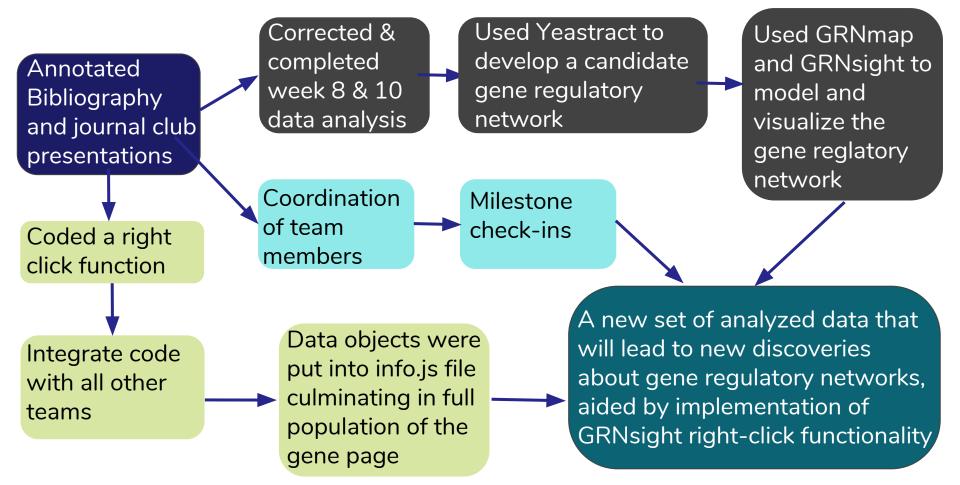


The right click functionality will be very useful for gene research

- Current Functionality: Visualize gene regulatory networks
- Goal: ability to easily open gene information page
- Solution: Adding right click functionality
 - Readily available gene information
 - No need for extensive database searching

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Individual milestones contribute to group effort



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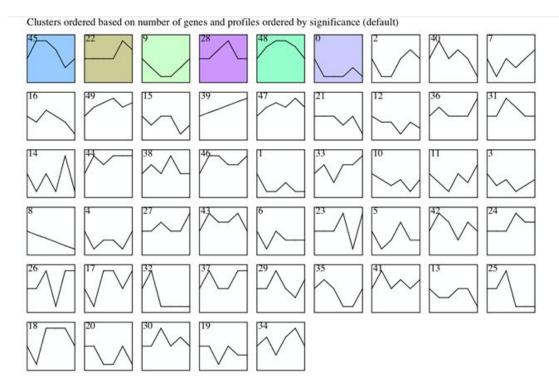
Percentage of genes demonstrating a cold shock response decreased with stricter p-values

- Standard p < 0.05 revealed ~41% of genes demonstrating significant response to cold shock
- B & H corrected p < 0.05 revealed second largest percentage
- Bonferroni-corrected p < 0.05 was the most strict

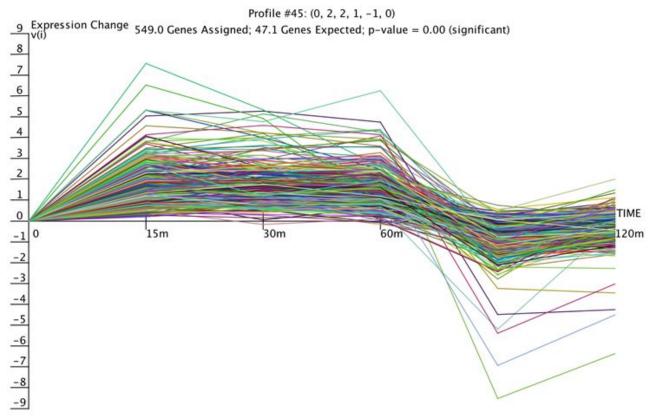
| ANOVA | WT |
|---|---------------|
| p < 0.05 | 2528 (40.85%) |
| p < 0.01 | 1652 (26.70%) |
| p < 0.001 | 919 (14.85%) |
| p < 0.0001 | 496 (8.01%) |
| Benjamini & Hochberg- corrected p < 0.05 | 1822 (29.44%) |
| Bonferroni-corrected p < 0.05 | 248 (4.01%) |

Genes were clustered into groups with similar regulation mechanisms

- Clusters ordered by significance
- Cluster 45 and 48 genes demonstrate upregulation
- Cluster 9 and 0 demonstrate downregulation



Profile 45 demonstrates clear upregulation after cold shock



A table of GO terms sheds light on the function of the regulatory genes

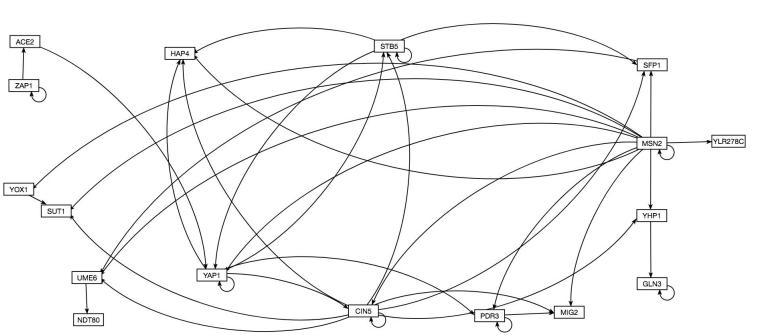
| Category Name | #Genes Category | #Genes Assigned | #Genes Expected | #Genes Enriched | p-value | Corrected p-value | Fold |
|---|--------------------|--------------------|--------------------|--------------------|----------|----------------------|------|
| cellular component organization (GO:0016043) | 540 | 161 | 162.7 | -1.7 | 0.6 | 1 | 1 |
| nuclear lumen (GO:0031981) | 327 | 183 | 98.5 | 84.5 | 2.40E-27 | <0.001 | 1.9 |
| RNA binding (GO:0003723) | 215 | 116 | 64.8 | 51.2 | 6.30E-15 | <0.001 | 1.8 |

Transcription factors for the network were selected based on significance

| TF | NDT80 | YAP1 | PDR3 | UME6 | MIG2 | ZAP1 | HAP4 | CIN5 |
|---------|----------|--------------|--------------|--------------|--------------|--------------|--------------|-------|
| P-value | 9.74E-10 | 1.31E- 09 | 2.82E- 09 | 6.12E- 09 | 1.96E- 08 | 2.99E-0 3 | 3.02E- 02 | 0.806 |

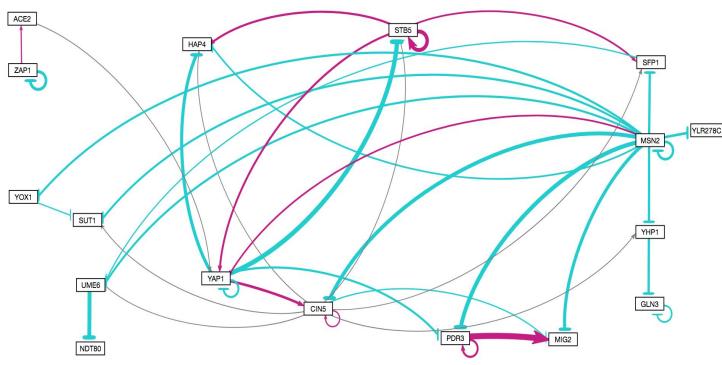
| TF | YOX1 | SFP1 | YHP1 | SUT1 | STB5 | ACE1 | MSN2 | GLN3 |
|---------|------|------|------|-------|-------|---------|----------|----------|
| P-value | 0 | 0 | 0 | 7E-15 | 7E-15 | 1.9E-14 | 2.72E-13 | 9.15E-11 |

Some genes are interconnected more than others



Arrows indicate interaction present

A weighted network details the nature of the interaction



- Pink lines indicate induction
- Blue lines
 indicate
 repression
- Thickness of lines indicate magnitude of interaction

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Right click functionality allows for easy access to additional gene information

```
.on("dblclick", nodeTextDblclick)
.on("contextmenu", function (gene) {
   var tempLink = $("<a></a>")
        .attr({
            href: "/gene/info.html?" + $.param({symbol: gene.name}),
            target: "_blank"
        });
   $("body").append(tempLink);
    tempLink.get(0).click();
    tempLink.remove();
   d3.event.preventDefault();
```



Inter team collaboration aided in development of new functionality

Web app and service for visualizing models of gene regulatory networks

S000005111 etc. YNL167C <name xmIns="http://uniprot.org/uniprot">SKO1_YEAST</name> MA0382.1

SKO1Genus Species

GRNSight

| General Information |
|------------------------------------|
| DNA Sequence |
| Protein Information |
| Gene Map |
| Regulation |
| Interaction |
| Gene Ontology |
| Frequency Matrix and Sequence Logo |

Inter team collaboration aided in development of new functionality

var sgdHrefTemplate = "https://www.yeastgenome.org/locus/"; var sgdId = gene.sgd.sgdID; \$(".sgd-link").text(sgdId).attr({ href: sgdHrefTemplate + sgdId });

// DNA Sequence Tab
var ensemblDNA = gene.ensembl.dnaSequence;
\$(".dnaSequence").text(ensemblDNA).attr({ href: ensemblHrefTemplate + ensemblDNA });

Frequency Matrix and Sequence Logo help visualize data from JASPAR



 A:0
 0
 0
 4921

 C:0
 0
 100841518

 G:1001000
 163728

 T:0
 0
 0
 0
 32

Code for Frequency Matrix and Sequence Logo

var frequencyMatrix = gene.jaspar.frequencyMatrix; var a = ""; for (var i = 0; i < frequencyMatrix.A.length; i++) { a += "" + frequencyMatrix.A[i] + ""; }

var sequenceLogo = gene.jaspar.sequenceLogo;
\$(".sequenceLogo").attr({ src : sequenceLogo });

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This team is confident that our contributions will aid future research

- Studying wild type yeast can show importance of certain genes in cold shock response when compared to mutated cells.
- Dynamic integration enhances user interface by providing the user with in depth information on the desired gene.
- Newly analyzed data and newly coded functionality will both lead to new discoveries about gene regulatory networks.

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- Microarray data of cold shock response in other eukaryotes
- Microarray data of other stress responses in yeast
- Prediction of related genes once right click function has been executed
- A function on the new gene page to alert the user that the page is loading or that data is being pulled

Summary

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