ALDH2 and CVD

Group M

ALDH2 and CVD

How does a decrease in function of the ALDH2 enzyme contribute to the development of cardiovascular diseases (CVD) such as coronary artery disease?

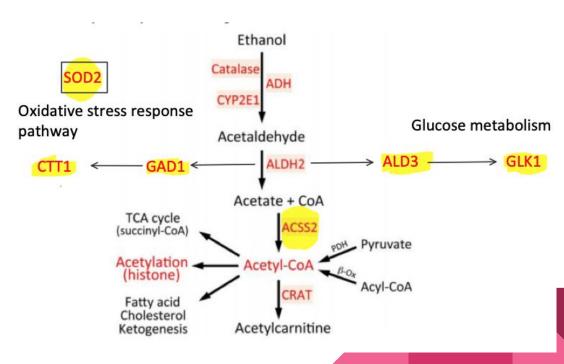
- ALDH2 → acetaldehyde and 4-HNE
- Other biomarkers?

Applying mathematical results

 Abstract results from graph theory (community detection and tests for centrality) applied to the PPIN of brewers yeast

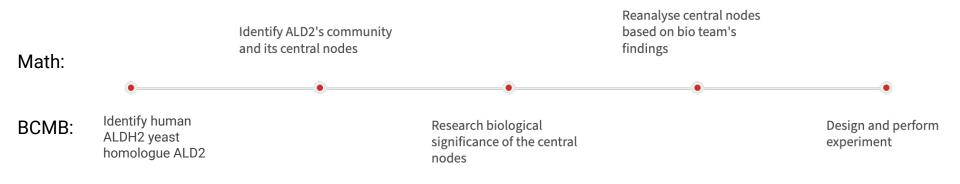
Proteins of interest

- •ACSS2
- •SOD2
- •GAD1
- •CTT1
- •GLK1
- •ALD3
- •Highly conserved pathway except for CCT1.

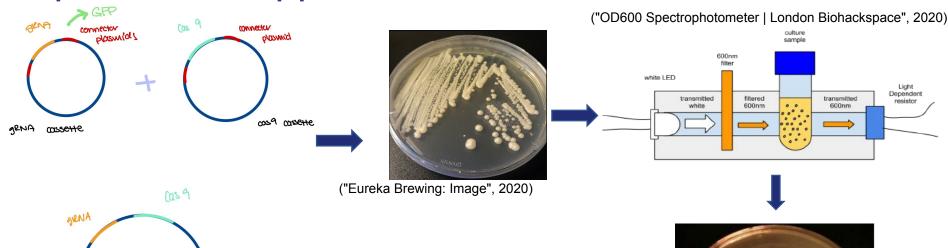


(Bruner, 2016)

Integration of BCMB and MATHS



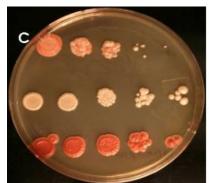
Experimental approaches



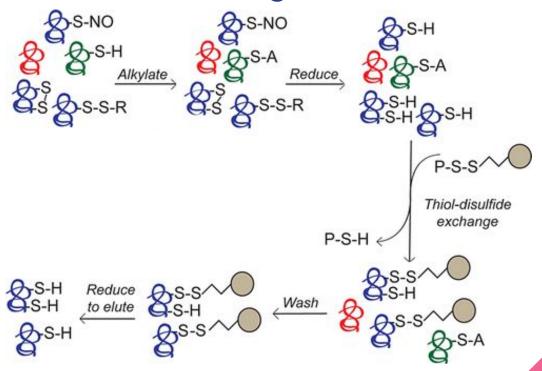
SELF-CONTAINED
COUSSE PLAS HID

(Akhmetov et al., 2018)

- Generation of a self-contained CRISPR plasmid
- Yeast culture, spectrophotometry and spot plate to assess viability during treatment of EtOH.

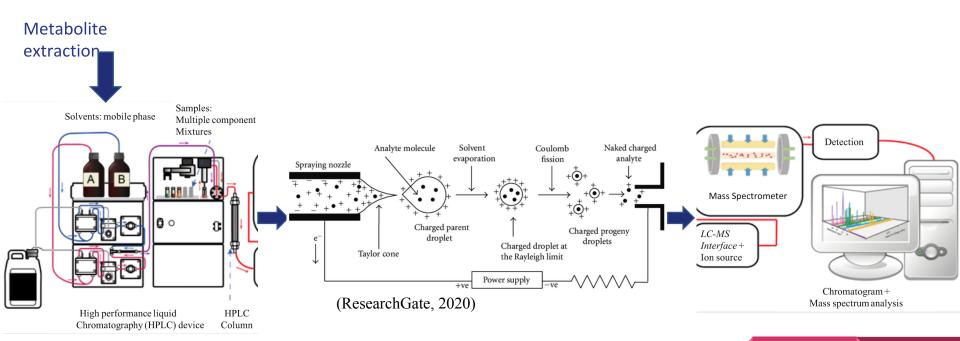


Thiol Disulfide Exchange



(Paulech et al., 2013)

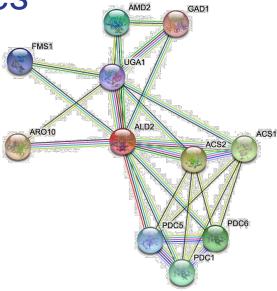
LCMS workflow



(Cronan, 2004)

Experimental approach: Mathematics

- Delete high degree nodes from the network
- Use Louvain community detection method to separate the graph into smaller communities
- Use centrality measures to identify central nodes within the community of ALD2
- Report central nodes to biochem team
- Discard central nodes which the biochem team report as insignificant
- Use bridges/shortest path to identify nodes on the path between ALD2 and the significant nodes



STRING interaction network (2020)

Conclusion

BCMB

 Comprehensive set of experimental procedures to investigate the importance of ALDH2 in regulating oxidative stress of and other biomarkers that may contribute to development of CVD

MATH

 The specific notions of "centrality" and "community" implied by each type of centrality measure and community detection method (respectively) need to be further investigated.

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