

	Gephebase Gene	GepheID
metallothionein CUP1 (https://www.gephebase.org/search-criteria?/and+Gene+Gephebase=^metallothionein CUP1^#gepbase-summary-title)	GP00001289	
	Entry Status	Arnoult
Published		Main curator

[illegible]

CUP1-1	Generic Gene Name	UniProtKB Saccharomyces cerevisiae (strain ATCC 204508 / S288c)
		P0CX80 (http://www.uniprot.org/uniprot/P0CX80)
CUP1; MTH1; YHR053C	Synonyms	GenebankID or UniProtKB
		856452 (https://www.ncbi.nlm.nih.gov/nuccore/856452)
4932.YHR055C	String	
(http://string-db.org/newstring.cgi/show_network_section.pl?identifier=4932.YHR055C)		
	Sequence Similarities	
Belongs to the metallothionein superfamily. Type 12 family.		
	GO - Molecular Function	
GO:0016209 : antioxidant activity (https://www.ebi.ac.uk/QuickGO/term/GO:0016209)		
GO:0005507 : copper ion binding (https://www.ebi.ac.uk/QuickGO/term/GO:0005507)		
GO:0046870 : cadmium ion binding (https://www.ebi.ac.uk/QuickGO/term/GO:0046870)		
GO:0004784 : superoxide dismutase activity		
(https://www.ebi.ac.uk/QuickGO/term/GO:0004784)		
	GO - Biological Process	

GO:0071585 : detoxification of cadmium ion
(<https://www.ebi.ac.uk/QuickGO/term/GO:0071585>)
GO:0010273 : detoxification of copper ion
(<https://www.ebi.ac.uk/QuickGO/term/GO:0010273>)
GO:0046688 : response to copper ion
(<https://www.ebi.ac.uk/QuickGO/term/GO:0046688>)
GO:0019430 : removal of superoxide radicals
(<https://www.ebi.ac.uk/QuickGO/term/GO:0019430>)

GO - Cellular Component

GO:0005829 : cytosol (<https://www.ebi.ac.uk/QuickGO/term/GO:0005829>)

Presumptive Null

No ([https://www.gephebase.org/search-criteria?/and+Presumptive Null=~No~#gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Presumptive+Null=~No~#gephebase-summary-title))

Molecular Type

Gene Amplification ([https://www.gephebase.org/search-criteria?/and+Molecular Type=~Gene Amplification~#gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Molecular+Type=~Gene+Amplification~#gephebase-summary-title))

Aberration Type

Complex Change ([https://www.gephebase.org/search-criteria?/and+Aberration Type=~Complex Change~#gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Aberration+Type=~Complex+Change~#gephebase-summary-title))

Molecular Details of the Mutation

Gene duplication

Experimental Evidence

Association Mapping ([https://www.gephebase.org/search-criteria?/and+Experimental Evidence=~Association Mapping~#gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Experimental+Evidence=~Association+Mapping~#gephebase-summary-title))

Main Reference

Too much of a good thing: the unique and repeated paths toward copper adaptation. (2015) (<https://pubmed.ncbi.nlm.nih.gov/25519894>)

Authors

Gerstein AC; Ono J; Lo DS; Campbell ML; Kuzmin A; Otto SP

Abstract

Copper is a micronutrient essential for growth due to its role as a cofactor in enzymes involved in respiration, defense against oxidative damage, and iron uptake. Yet too much of a good thing can be lethal, and yeast cells typically do not have tolerance to copper levels much beyond the concentration in their ancestral environment. Here, we report a short-term evolutionary study of *Saccharomyces cerevisiae* exposed to levels of copper sulfate that are inhibitory to the initial strain. We isolated and identified adaptive mutations soon after they arose, reducing the number of neutral mutations, to determine the first genetic steps that yeast take when adapting to copper. We analyzed 34 such strains through whole-genome sequencing and by assaying fitness within different environments; we also isolated a subset of mutations through tetrad analysis of four lines. We identified a multilayered evolutionary response. In total, 57 single base-pair mutations were identified across the 34 lines. In addition, gene amplification of the copper metallothionein protein, CUP1-1, was rampant, as was chromosomal aneuploidy. Four other genes received multiple, independent mutations in different lines (the vacuolar transporter genes VTC1 and VTC4; the plasma membrane H⁺-ATPase PMA1; and MAM3, a protein required for normal mitochondrial morphology). Analyses indicated that mutations in all four genes, as well as CUP1-1 copy number, contributed significantly to explaining variation in copper tolerance. Our study thus finds that evolution takes both common and less trodden pathways toward evolving tolerance to an essential, but highly toxic, micronutrient.

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Additional References

RELATED GEPHE

Related Genes

5 (MAM3, Plasma membrane ATPase 1, PMR1, Vacuolar transporter chaperone 1, Vacuolar transporter chaperone 4) ([https://www.gephebase.org/search-criteria?/or+Taxon ID=~4932~/and+Trait=Metal tolerance/and+groupHaplotypes=true#gephebase-summary-title](https://www.gephebase.org/search-criteria?/or+Taxon+ID=~4932~/and+Trait=Metal+tolerance/and+groupHaplotypes=true#gephebase-summary-title))

Related Haplotypes

No matches found.

EXTERNAL LINKS

COMMENTS

24 over 34 lines