

*Patterns of genetic diversity and
reproductive isolation in natural
Saccharomyces yeast populations.*

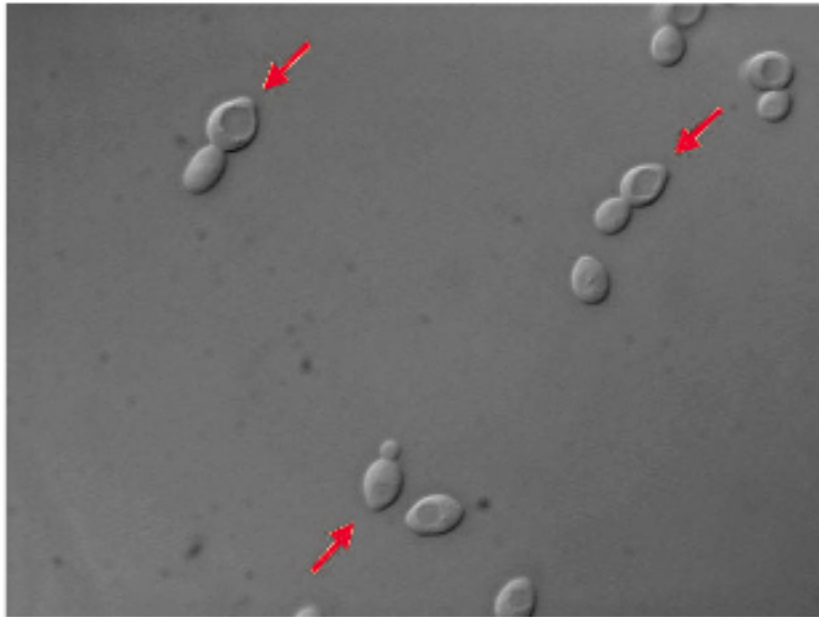
KITP, Santa Barbara
November 2008

Paul Sniegowski, Department of Biology, University of
Pennsylvania

Overview

- Who is *Saccharomyces cerevisiae* and how is it related to its congeners?
- The evolutionary and ecological status of natural *Saccharomyces* populations.
- Genetic structure of natural *Saccharomyces* populations.
- Patterns of reproductive isolation in *S. paradoxus*.
- Conclusion and future prospects.

Unicellular Ascomycete Fungus



Asexual Budding



Four Spores in an Ascus



Saccharomyces cerevisiae



S. cerevisiae

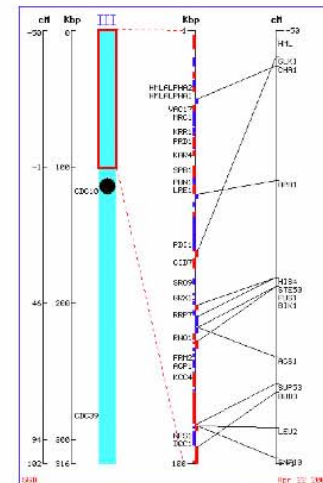
Genetics

Cell and Molecular Biology

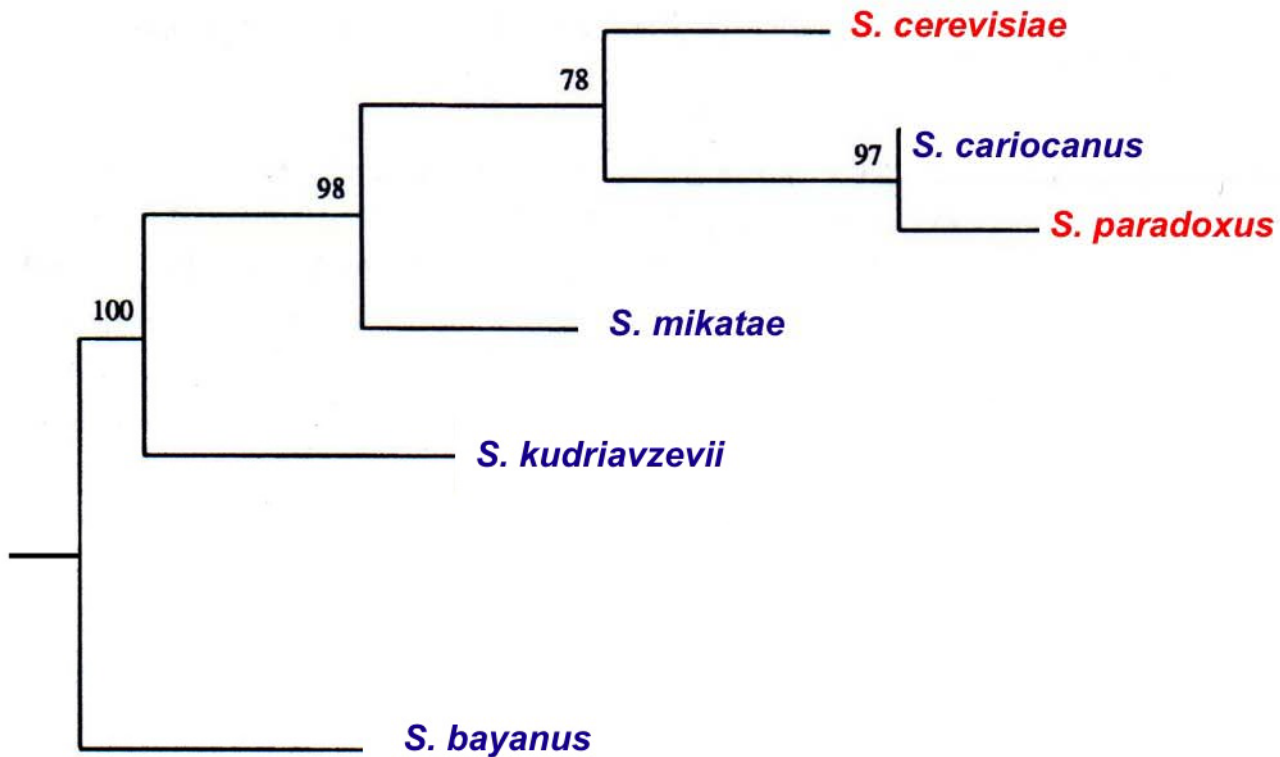
Proteomics

Genomics

Ecology and Evolution



Saccharomyces sensu stricto



After Naumov et al. 2000

Yeast in Nature



Woodland Contexts

Yeast Field Collection



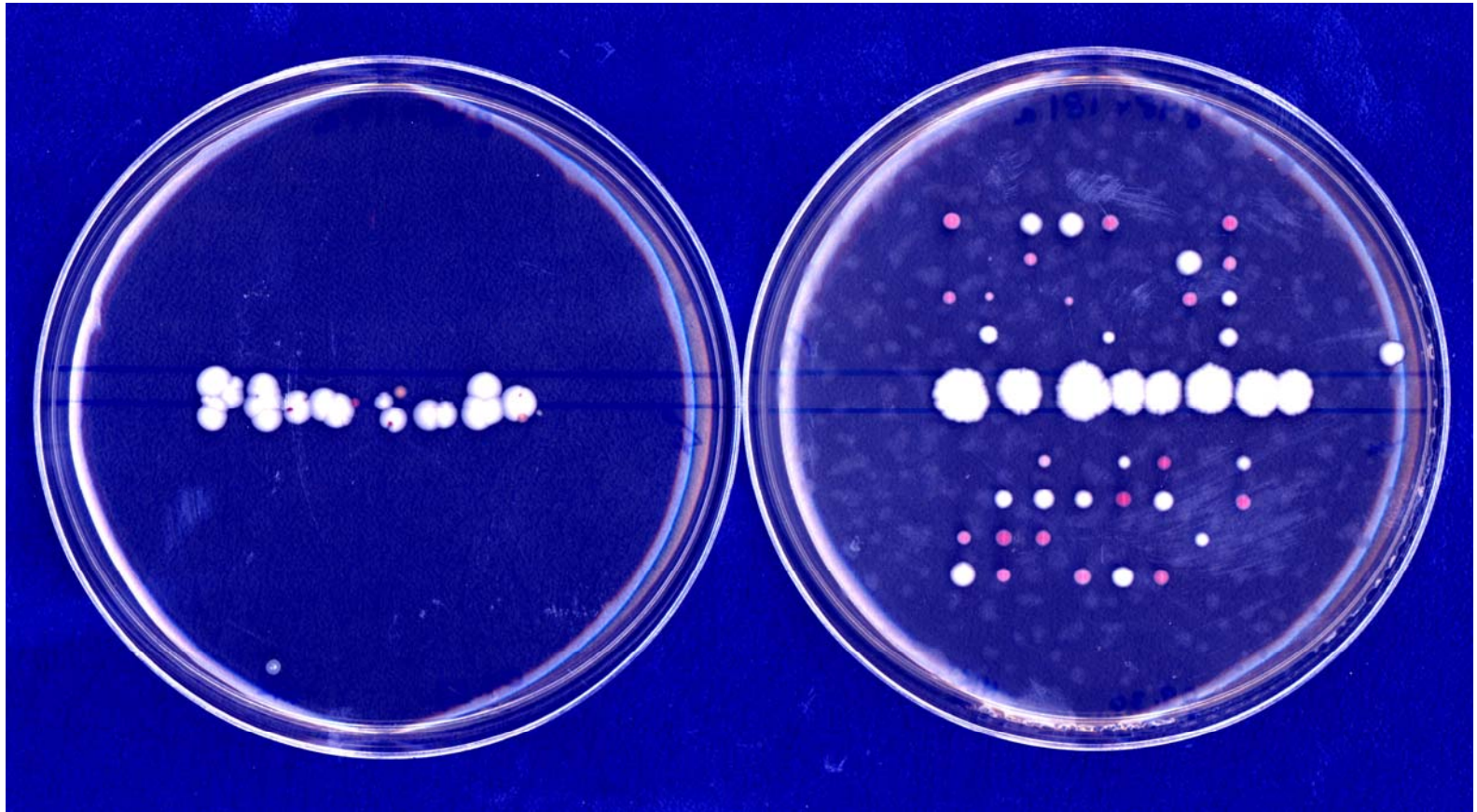
Sampling oak exudate in Ontario, Canada

Yeast Isolation



Laboratory Contexts

Testing for reproductive isolation



Heterospecific test cross

Conspecific test cross

***Saccharomyces paradoxus* and *Saccharomyces cerevisiae* are associated with exudates of North American oaks**

Gennadi I. Naumov, Elena S. Naumova, and Paul D. Sniegowski

Table 1. *Saccharomyces* strains from which monosporic cultures were used.

Species	Strain	Source	Place of isolation	Author
<i>S. cerevisiae</i>	VKM Y-502	Grape berries	Russian Far East	I.A. Mazilkin
	YNN 295	Genetic line	—	D. Vollrath and R.W. Davis
	95-5	<i>Quercus</i> sp.	Long Woods, W.K. Kellogg Biological station, Michigan State Univ.	G.I. Naumov
	95-6	<i>Quercus</i> sp.	Long Woods, W.K. Kellogg Biological station, Michigan State Univ.	G.I. Naumov
<i>S. paradoxus</i>	CBS 5829	Mor soil	Denmark	V. Jensen
	CBS 432	Unknown	Unknown	Unknown
	95-1	<i>Quercus</i> sp.	Saugatuck Dunes State Park, Mich.	G.I. Naumov
	95-3	<i>Q. alba</i>	Long Woods, W.K. Kellogg Biological station, Michigan State Univ.	G.I. Naumov
	95-4	<i>Q. rubra</i>	Long Woods, W.K. Kellogg Biological station, Michigan State Univ.	G.I. Naumov
	95-7	<i>Q. rubra</i>	Michigan State Univ. Campus, East Lansing	G.I. Naumov
	95-8	<i>Q. rubra</i>	Michigan State Univ. Campus, East Lansing	G.I. Naumov
	52-153 ^a	<i>Drosophila</i> sp.	U.S.A.	H.J. Phaff
	61-359 ^a	<i>Ulmus</i> sp.	U.S.A.	H.J. Phaff
<i>Saccharomyces</i> sp.	97-3 ^b	<i>Q. alba</i>	Allaire State Park, N.J.	G.I. Naumov

Saccharomyces cerevisiae and *Saccharomyces paradoxus* coexist in a natural woodland site in North America and display different levels of reproductive isolation from European conspecifics

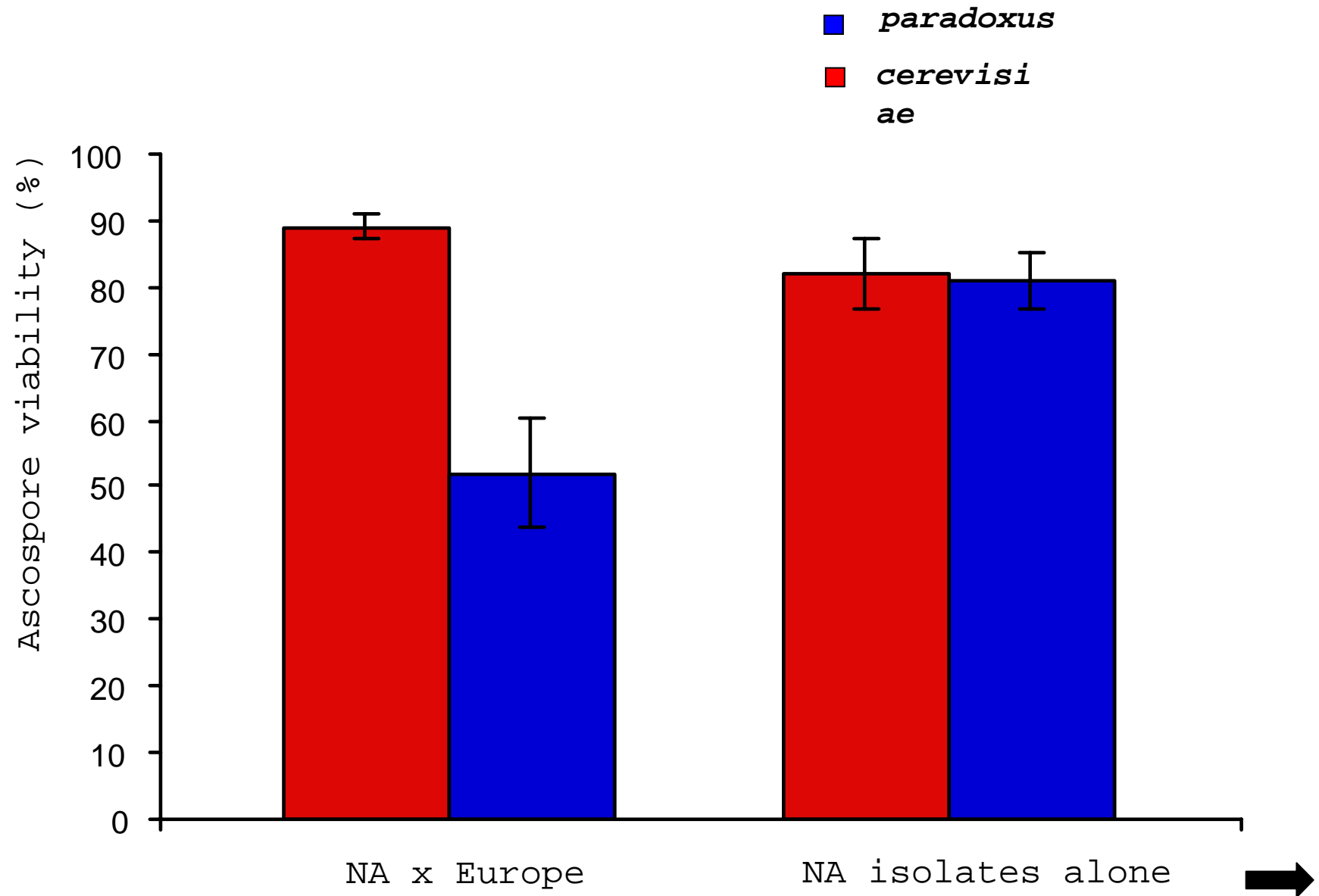
Paul D. Sniegowski *, Peter G. Dombrowski, Ethan Fingerman

Table 1

Identified isolates of *S. paradoxus* and *S. cerevisiae* and their source habitats in a Pennsylvanian woodland

Species	Strain	Source	
<i>S. cerevisiae</i>	YPS 128	Soil beneath <i>Q. alba</i>	
	YPS 129	Flux from <i>Q. alba</i>	
	YPS 133	Soil beneath <i>Q. alba</i>	
	YPS 134	Soil beneath <i>Q. velutina</i>	
	YPS 139	Soil beneath unidentified <i>Quercus</i> spp.	
	YPS 141	Soil beneath <i>Q. velutina</i>	
	YPS 142	Bark of <i>Q. rubra</i>	
	YPS 143	Soil beneath <i>Q. rubra</i>	
	YPS 154	Bark of <i>Q. velutina</i>	
	YPS 163	Soil beneath <i>Q. rubra</i>	
	<i>S. paradoxus</i>	YPS 125	Flux of <i>Q. rubra</i>
		YPS 138	Soil beneath <i>Q. velutina</i>
		YPS 145	Soil beneath <i>Q. alba</i>
YPS 150		Bark of <i>Q. velutina</i>	
YPS 151		Soil beneath <i>Q. velutina</i>	
YPS 152		Soil beneath <i>Q. rubra</i>	
YPS 155	Bark of <i>Q. rubra</i>		
YPS 158	Soil beneath <i>Q. alba</i>		

Geographic reproductive isolation in *paradoxus* but not *cerevisiae*?



Heidi A. Kuehne

Ph.D. Thesis, University
of Pennsylvania

The genetic structure
and biogeography
of natural *Saccharomyces*
populations.



Setting the Table

What is a population? Are there broad subdivisions in nature?

How are these populations structured genetically?

How are these populations structured spatially?

What types of breeding systems are realized in nature?



GENETIC POPULATION STRUCTURE

Mutation



Selection



GPS is the distribution and abundance of genetic variation within and among populations.

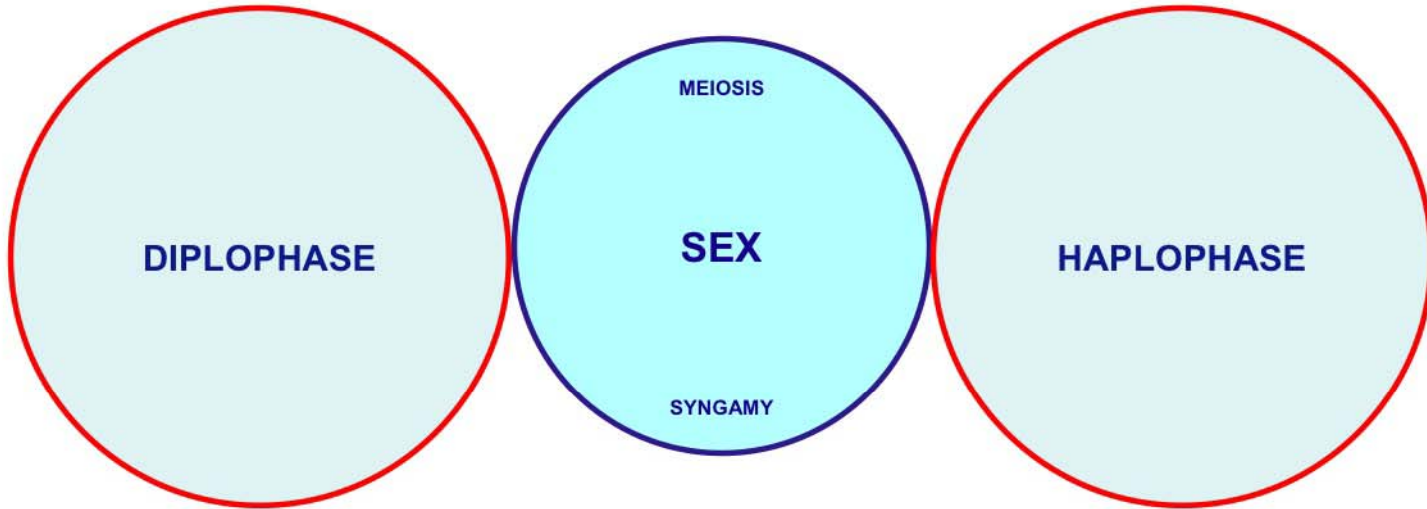
Migration



Drift

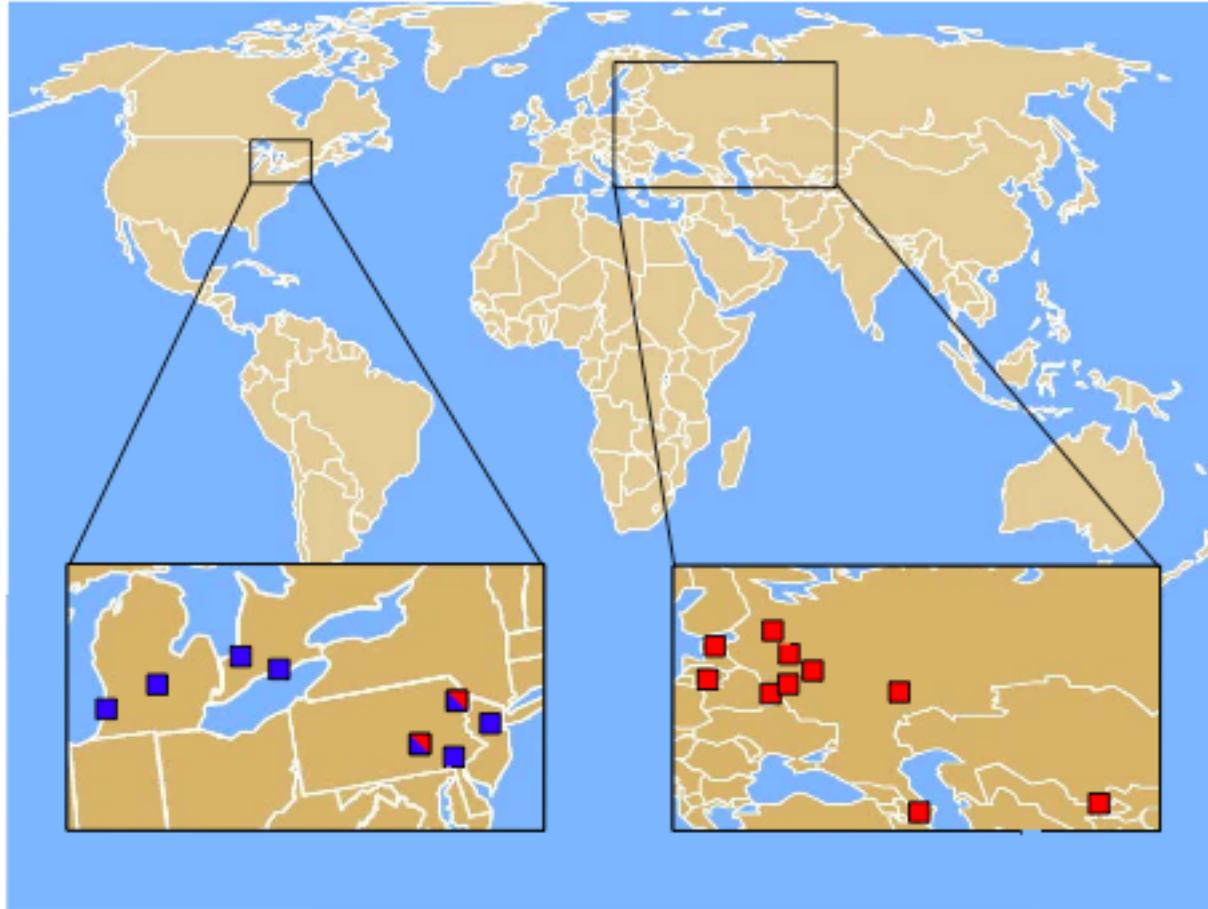


Haplodiplontic Life Cycle



Reproduction and Sex Are Uncoupled

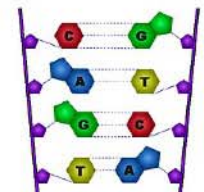
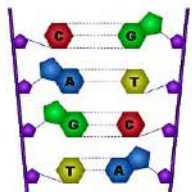
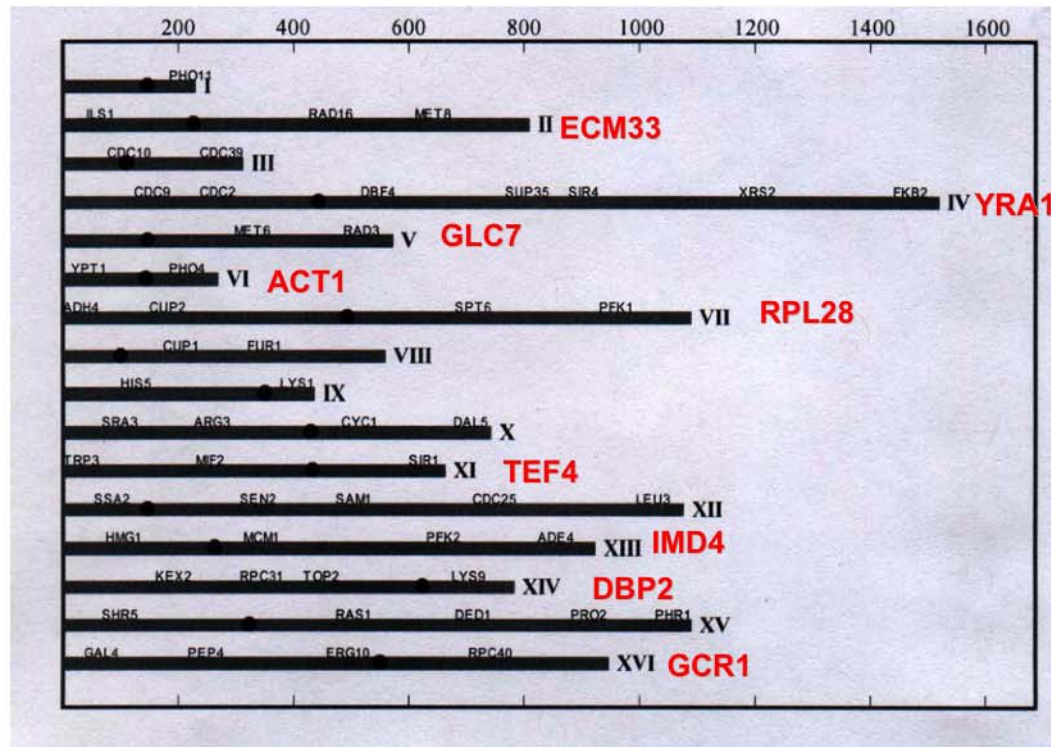
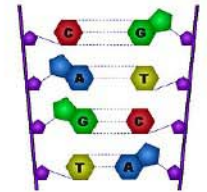
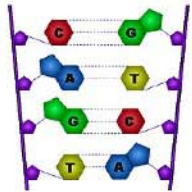
Yeast Collection Sites



51 *S. paradoxus*
27 *S. cerevisiae*

11 *S. paradoxus*

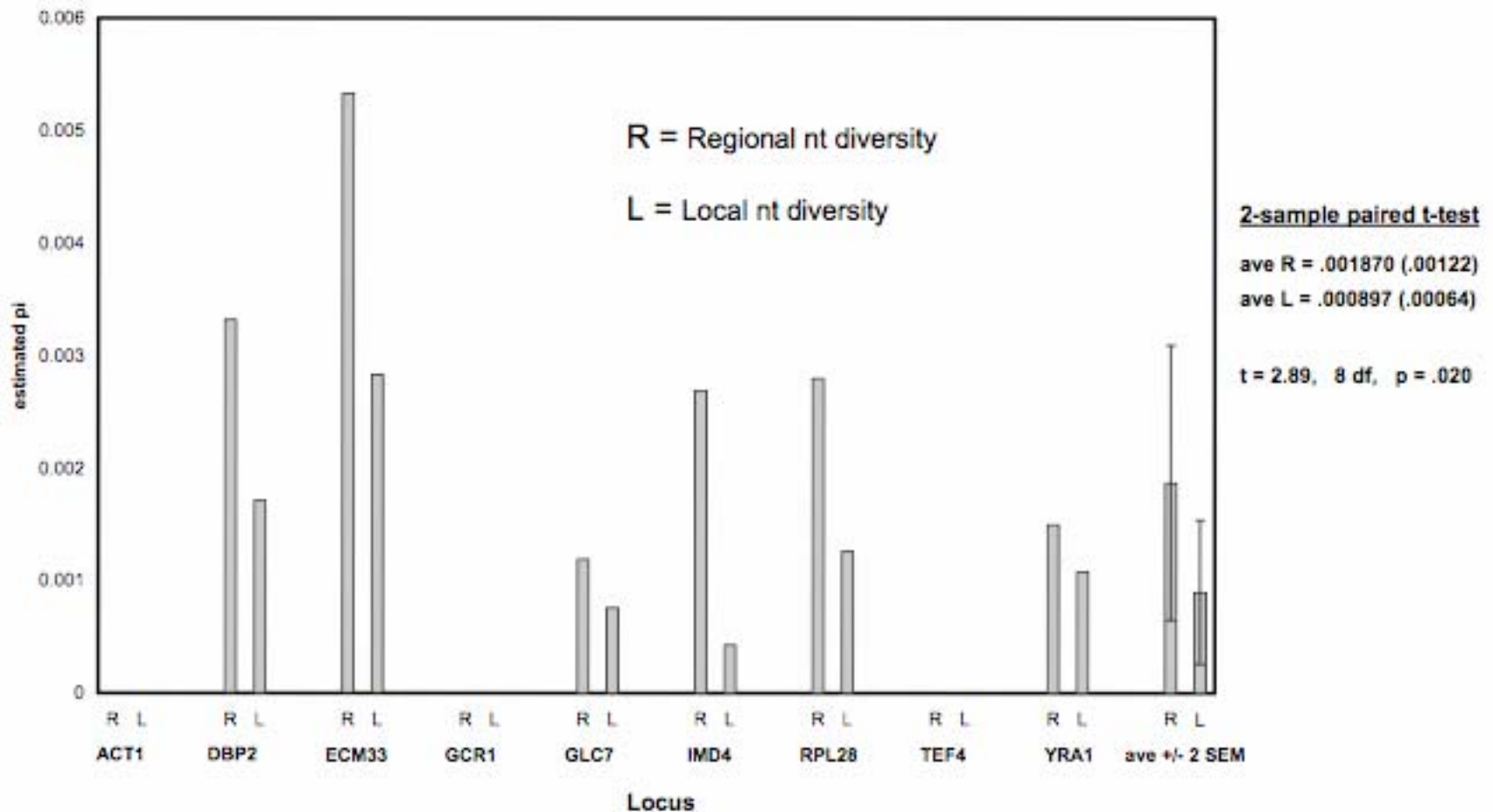
Multilocus Sequence Typing



Nine Spliceosomal Introns
350,000 nt

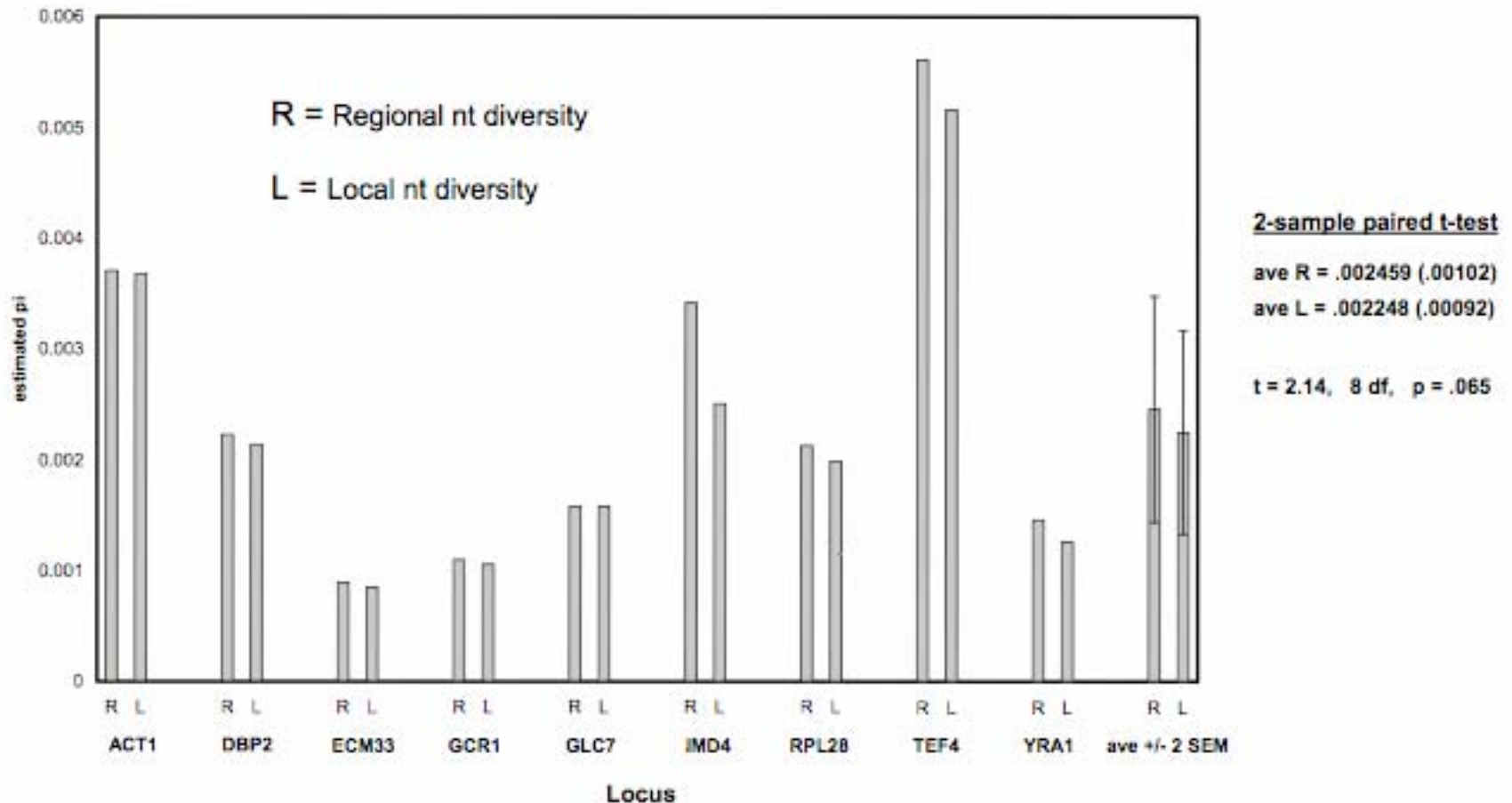
Diversity Patterns

S. cerevisiae Silent Nucleotide Diversity

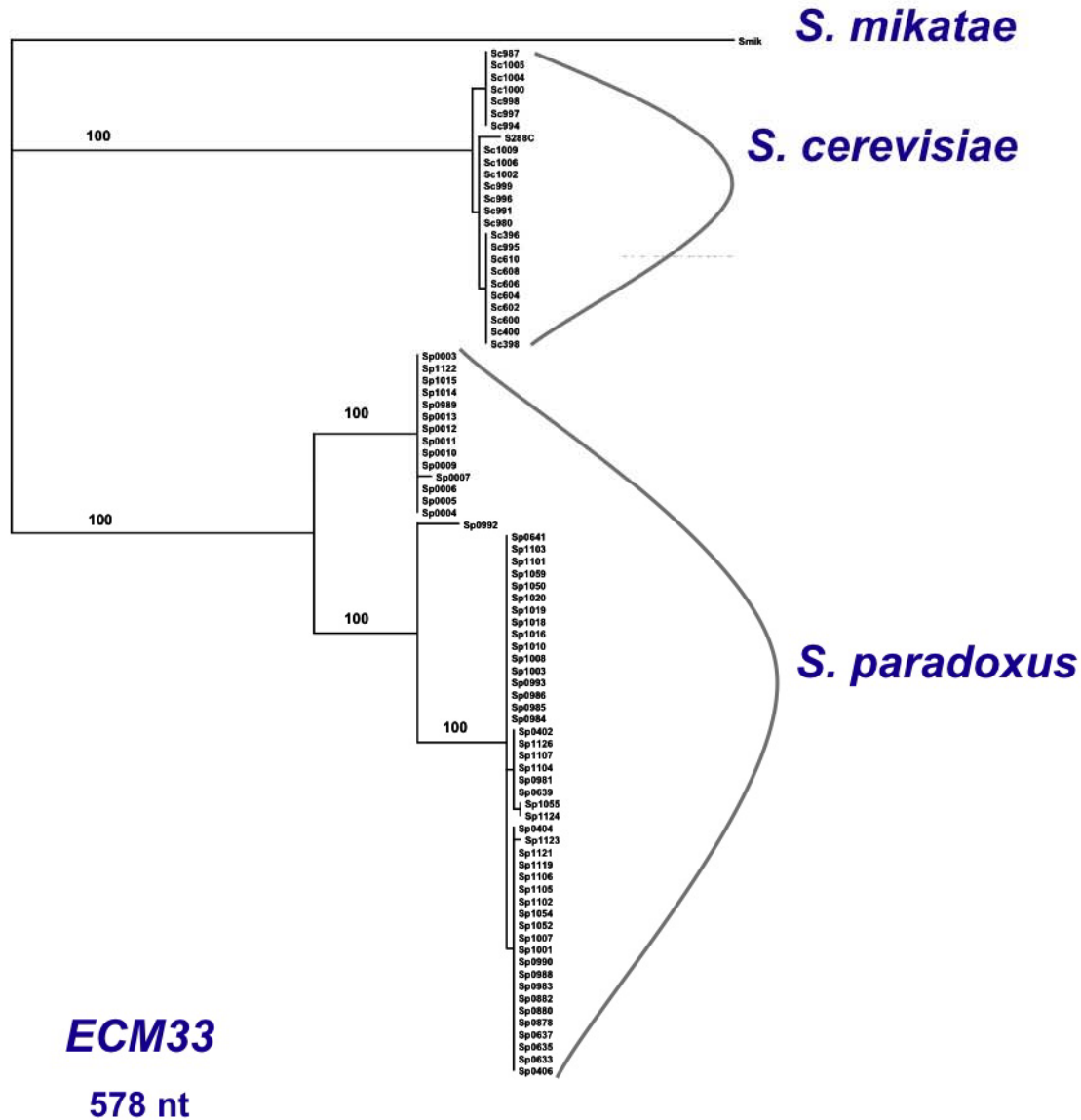


Diversity Patterns

S. paradoxus Silent Nucleotide Diversity



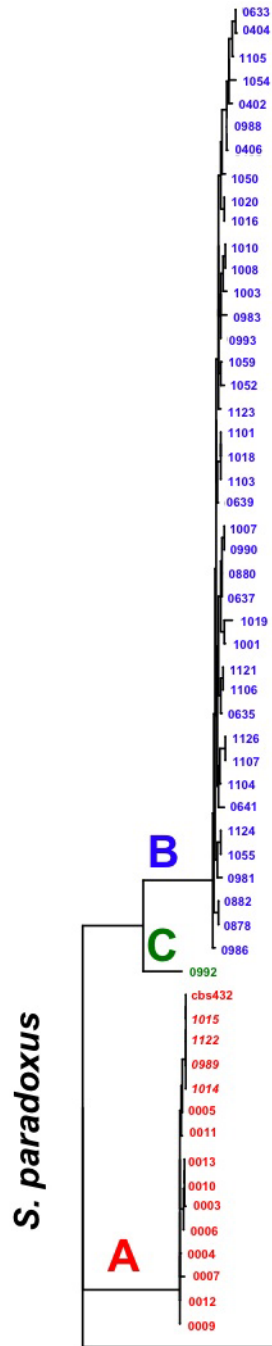
Maximum Parsimony Gene Trees



Distinct Genetic Groups

S. paradoxus

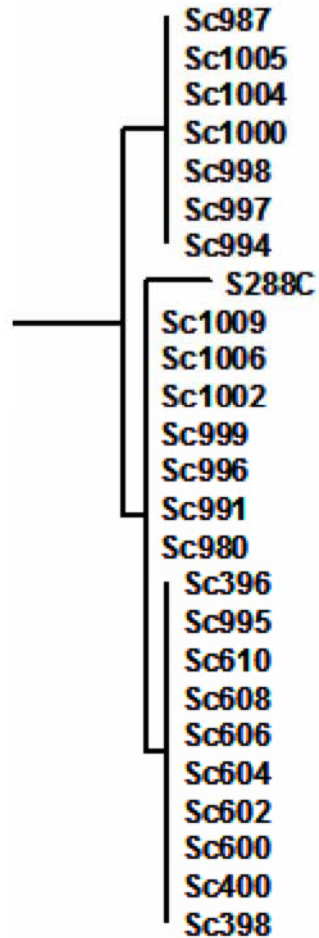
nine loci, 4867 bp



	F	S
Grps. A & B	179	0
Grps. A & C	165	0
Grps. B & C	89	0

Distinct Genetic Groups in Sympatry

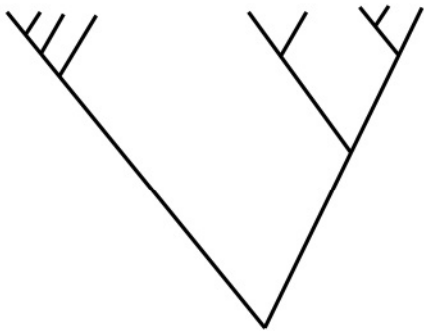
S. cerevisiae



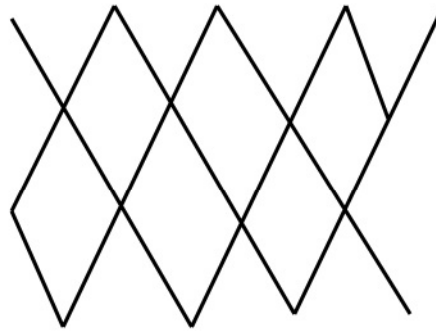
nine loci, 4839 bp

	F	S
<i>Grps. I & II</i>	6	0
<i>Grps. I & III</i>	15	0
<i>Grps. II & III</i>	11	0

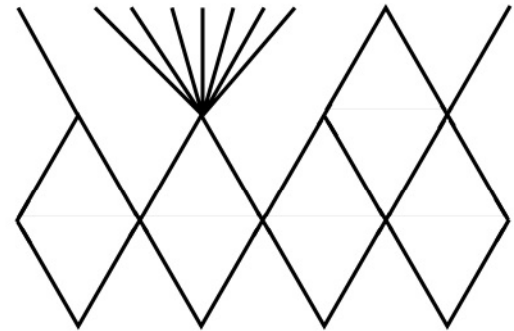
Possible Population Structures



Clonal



Recombining



Epidemic

After Maynard Smith *et al.* 1993

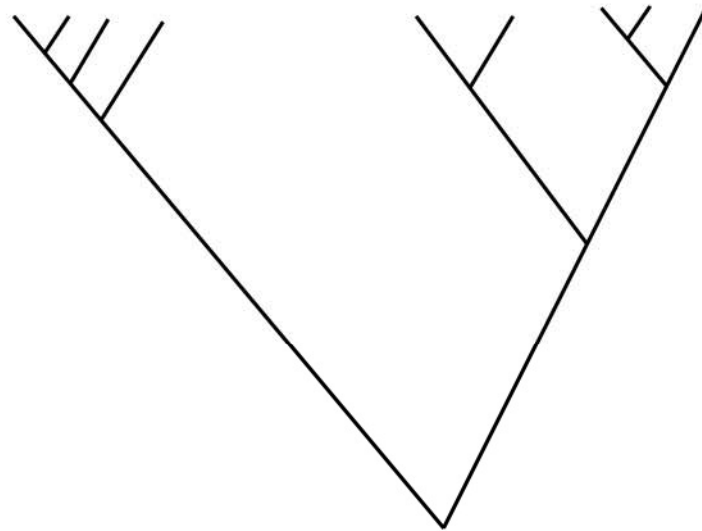
Testing population structure using multilocus sequence data

Index of Association (I_A): Tests for linkage disequilibrium, the nonrandom association of alleles in haplotypes, by analyzing the distribution of genetic distances.

Incongruency Length Difference (ILD): Tests whether different gene trees are congruent (clonality) or incongruent.

Consistency Index (CI): Compares trees for homoplasy (obs. character state arising more than once in tree by nonhomologous means). Recombination is a source of homoplasy.

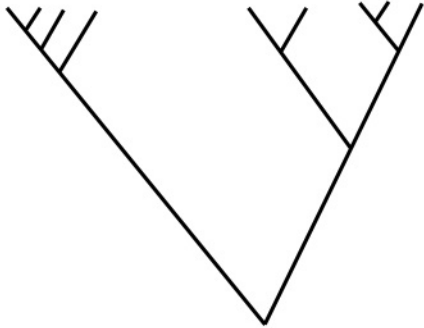
S. cerevisiae



I_A $p = .001$
ILD $p = 1.00$
CI = 1.00

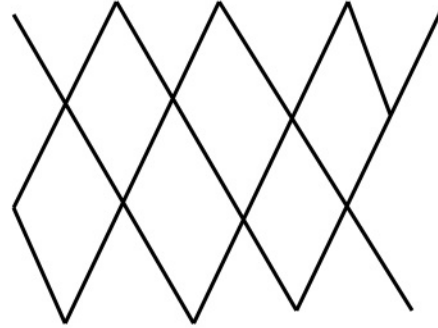
Clonal in Northeast American Woodlands

S. paradoxus



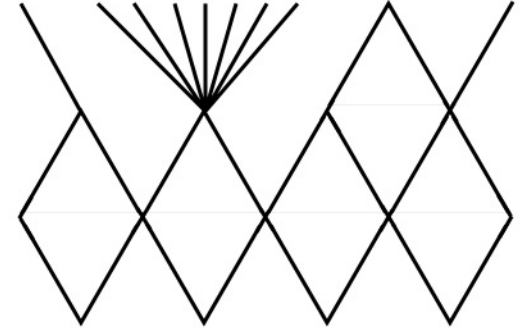
Group A
NE America

NA



Group A
Eurasia

I_A $p = .373$
ILD $p = .552$
CI = .86

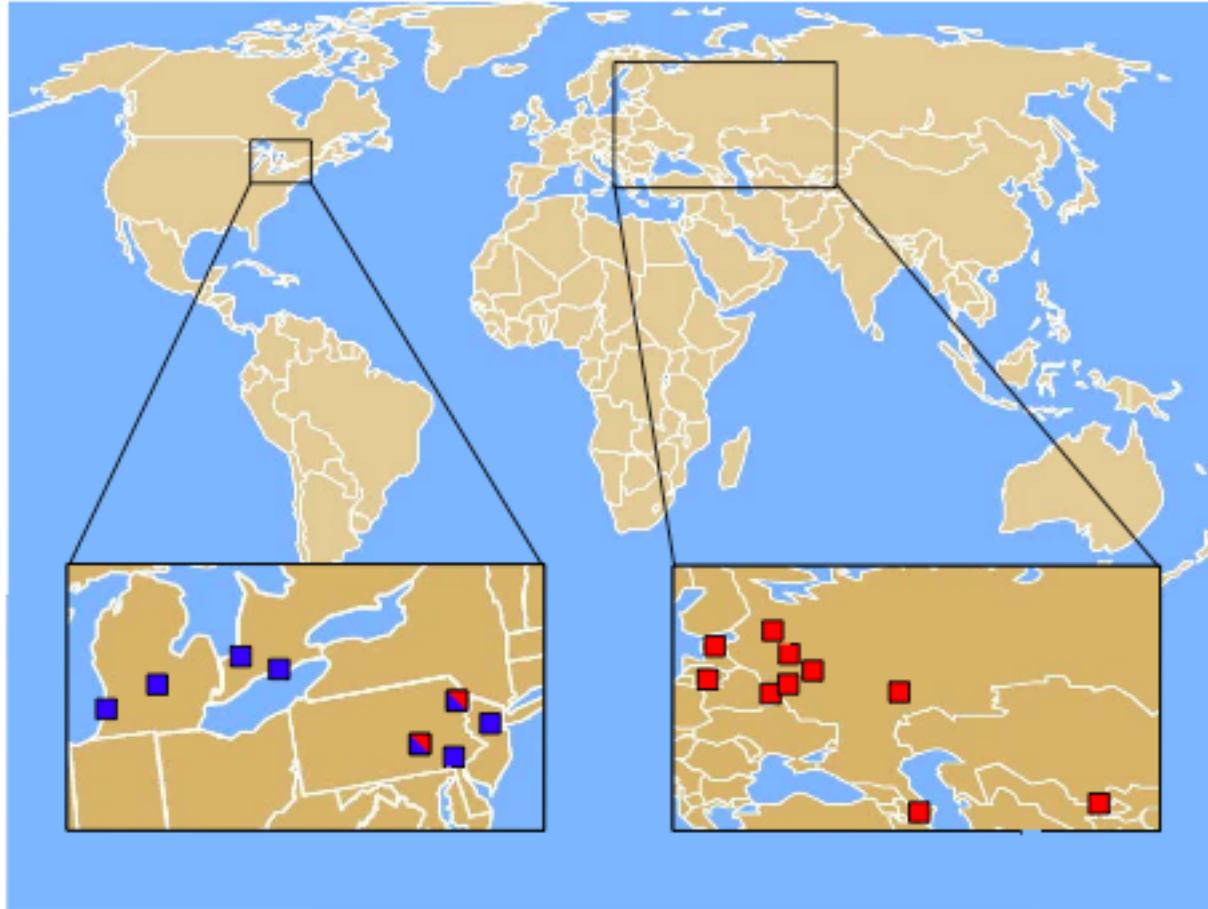


Group B
NE America

I_A $p = .1914$
ILD $p = .001$
CI = .41

Multiple Structures in the Northern Hemisphere

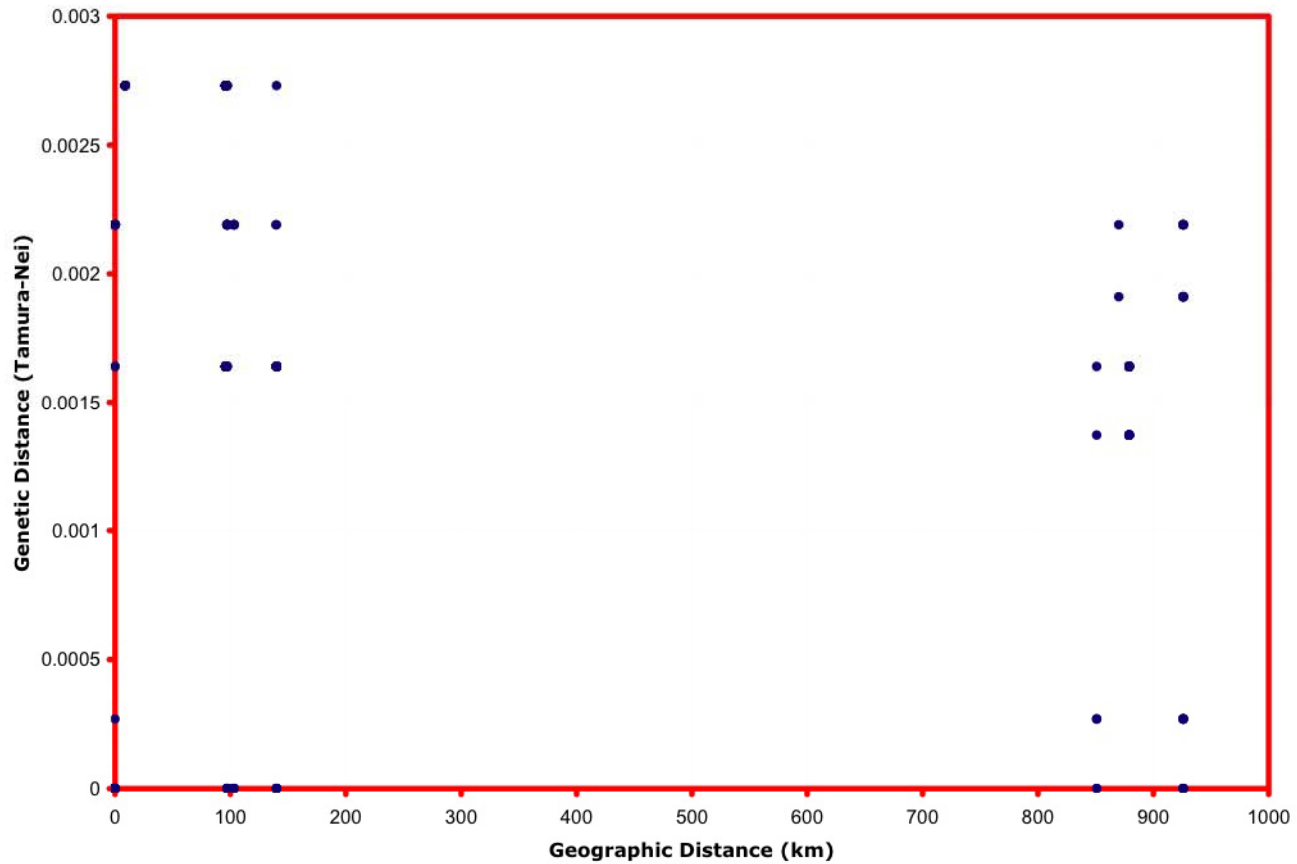
Yeast Collection Sites



51 *S. paradoxus*
27 *S. cerevisiae*

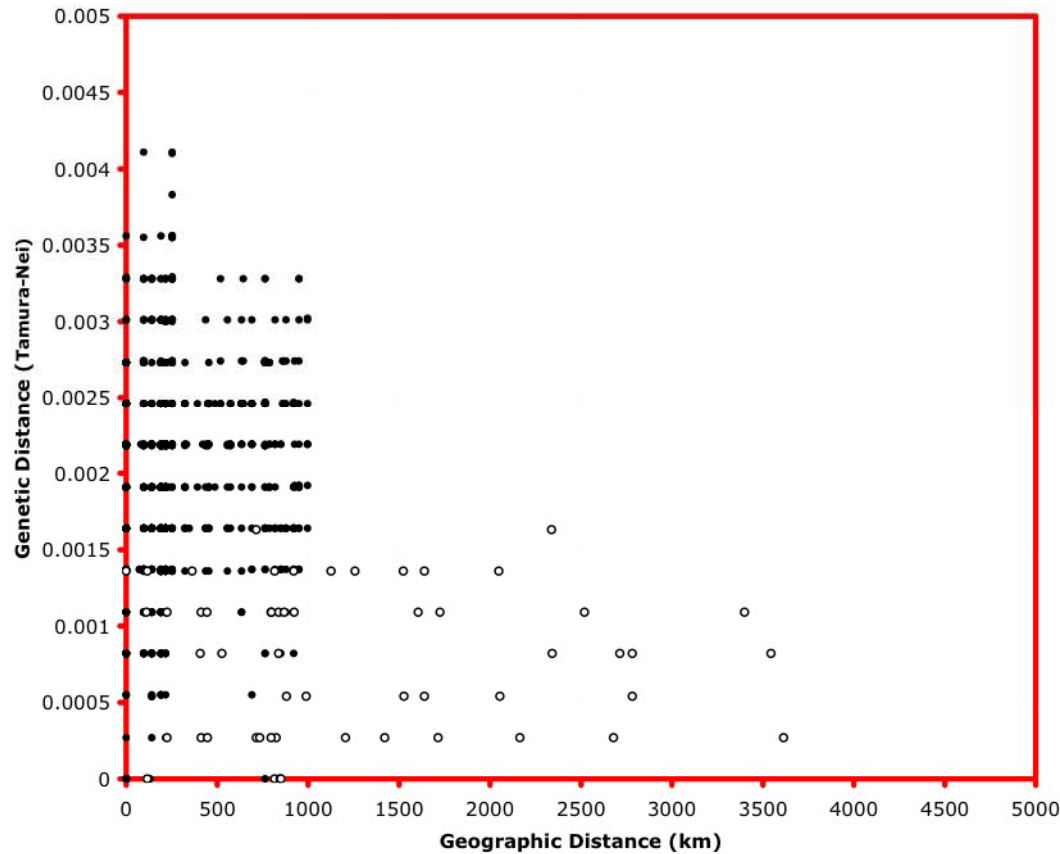
11 *S. paradoxus*

Regional Biogeography



S. cerevisiae

Regional Biogeography



Mantel Tests

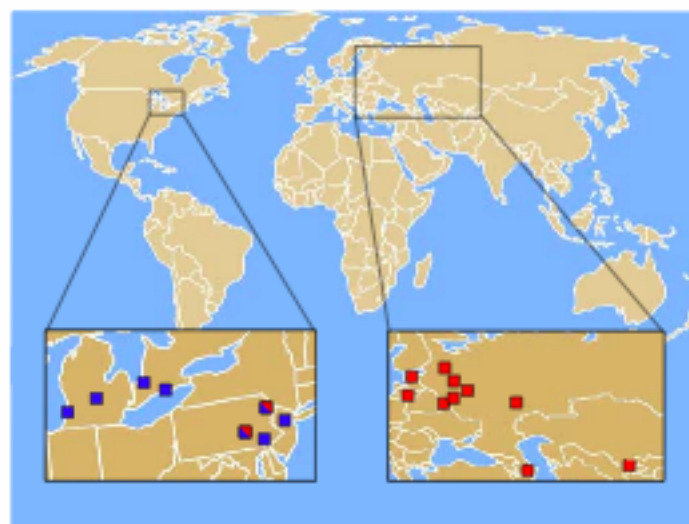
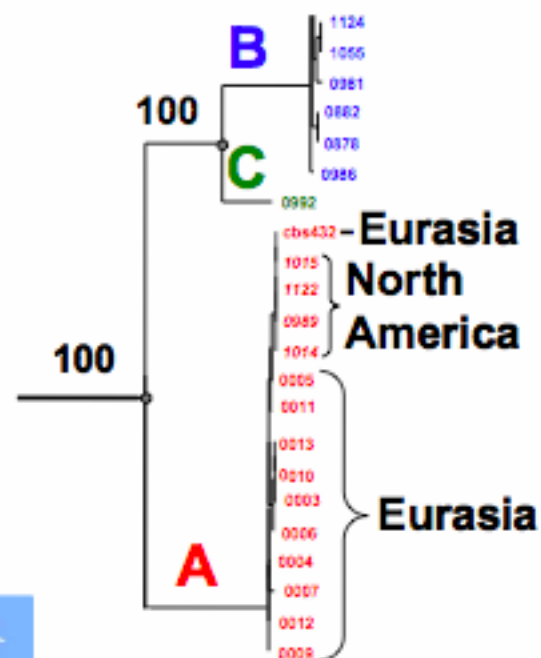
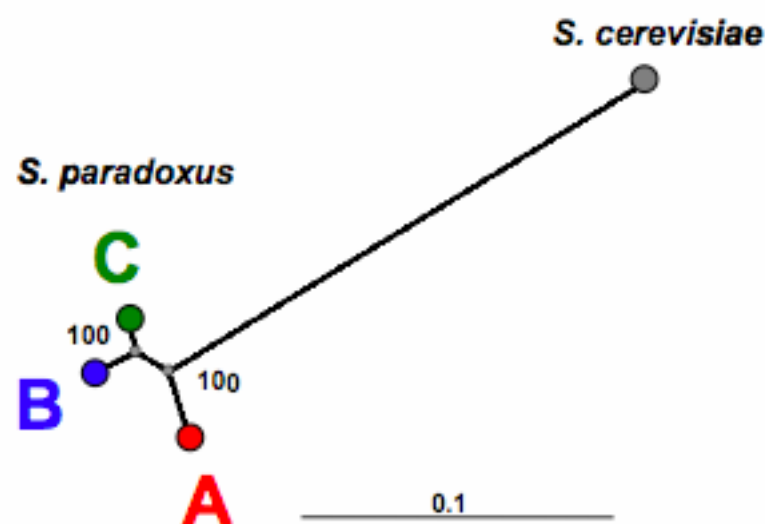
GRP. **B**(all) $p = .09$
(uni) $p = .22$

GRP. **A**(EE) $p = .45$

- Northeast America
- East Europe

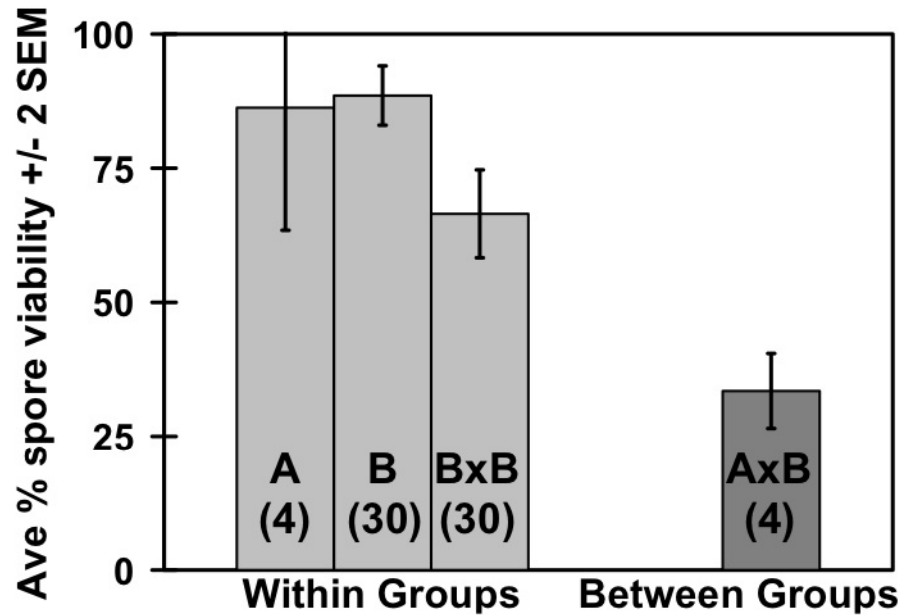
S. paradoxus

Global Biogeography: *S. paradoxus*



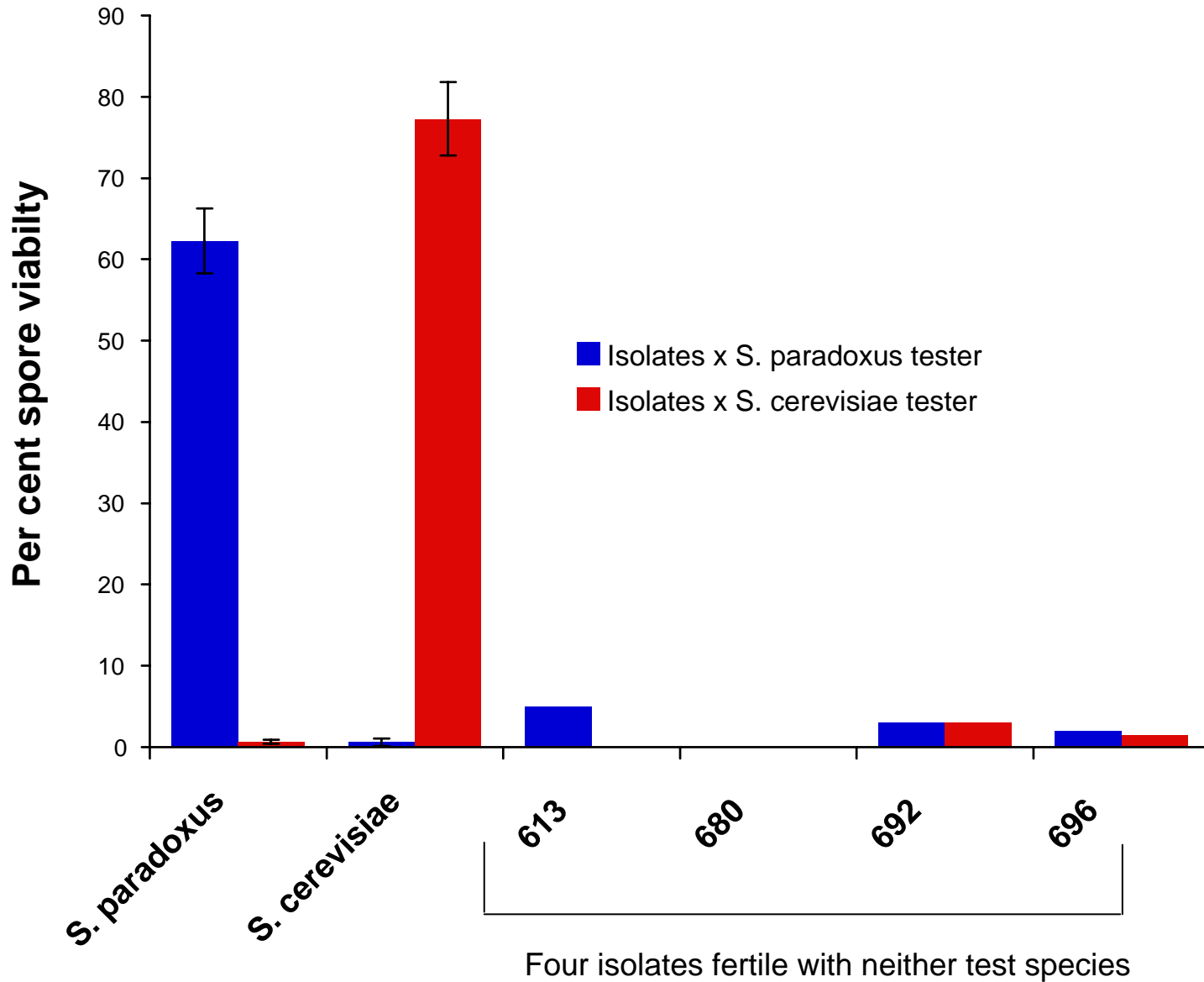
Evolution in allopatry followed by migration

Are These Cryptic Species?

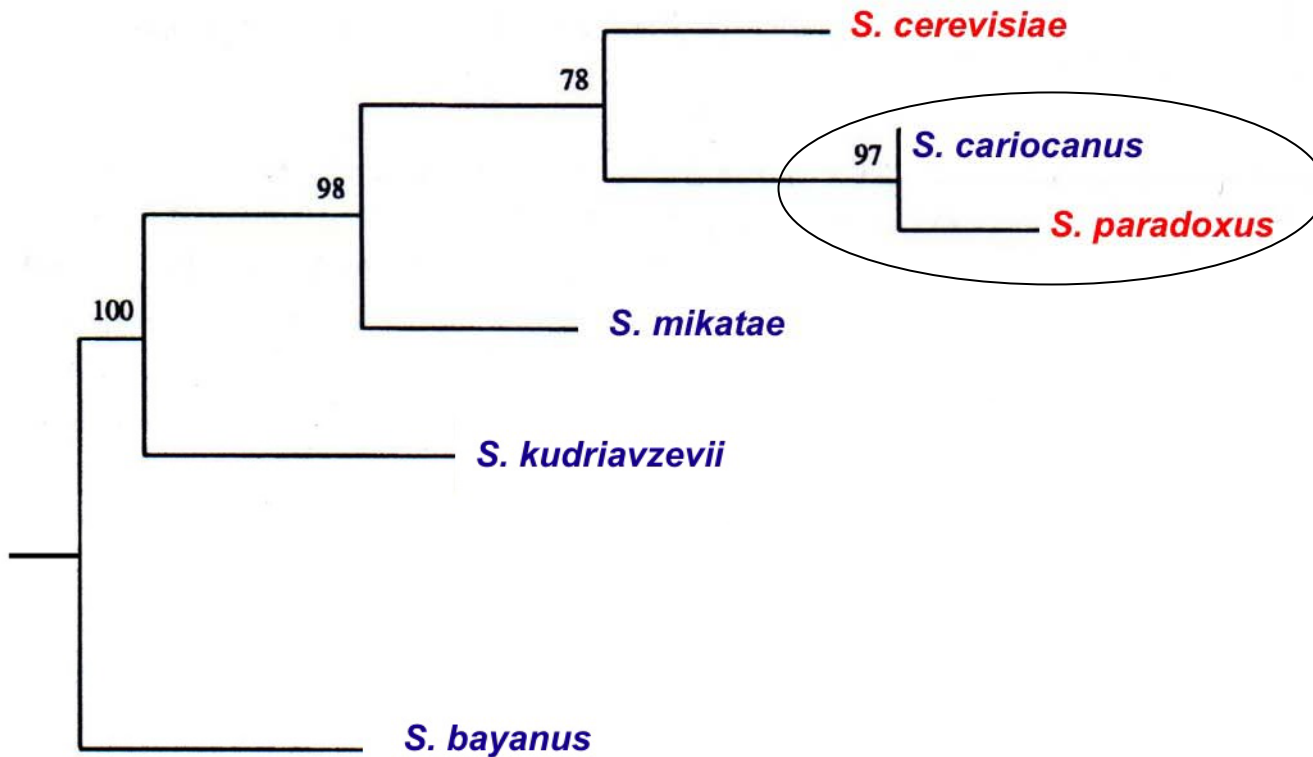


	A (N. America)	B (N. America)	C (N. America)
A (Eurasia)	3	176	160
A (N. America)		181	165
B (N. America)			89

"Mystery" isolates



Saccharomyces sensu stricto



After Naumov et al. 2000

What are the mystery isolates?

Three possibilities (assuming that they are all the same thing).

1. Hybrids. This appears to be ruled out by recent sequence data.
2. *S. cariocanus*. Sequence data can't rule this out, as *S. cariocanus* is *closer* in sequence to NA *S. paradoxus* (indeed, almost identical) than to EU *S. paradoxus*. (Interesting, huh?)
3. A newly diverging, previously unknown lineage closely related to *S. paradoxus*; a species *in statu nascendi*. (Most exciting to me...)

Conclusions

- Evidence suggests the existence of natural *S. cerevisiae* populations.
- S. cerevisiae* and its known wild congener *S. paradoxus* are sympatric in nature and occupy the same habitat, at least in North America.
- Within geographical region (e.g., NA, Eurasia) neither species shows much evidence of geographic population structure.
- The two species exhibit markedly different genetic population structures in nature: *cerevisiae* populations are clonal, *paradoxus* populations are recombining.
- paradoxus* exhibits genetic differentiation between continental regions and shows evidence of multiple, genetically isolated sympatric lineages, at least one of which is likely to have initially diverged in allopatry.
- There is emerging evidence (Duncan Greig, Helen Murphy) that prezygotic isolation and reinforcement are involved in speciation in yeast.

Some Future Prospects

--Does *S. cerevisiae* exhibit global population differentiation like *paradoxus*? (Are there globally distributed natural *cerevisiae* populations and are they genetically differentiated?)

--Nagging question: Are *S. cerevisiae* natural populations truly wild, or just feral?

--What is the nature of speciation processes in *Saccharomyces*? How many sympatric, genetically isolated lineages are there related to *S. paradoxus*, and what keeps them apart genetically? How big a role does mating behavior play in reproductive isolation? How important is allopatric divergence? Etc.

--What are the important ecological differences between natural *S. cerevisiae* and *paradoxus* populations?

--What are the important ecological, genomic and physiological differences within and between natural *Saccharomyces* populations?

Acknowledgments and Thanks

For actually doing much of the work described here: Heidi Kuehne (now postdoc U. Edinburgh) and Helen Murphy (now postdoc Wake Forest U.).

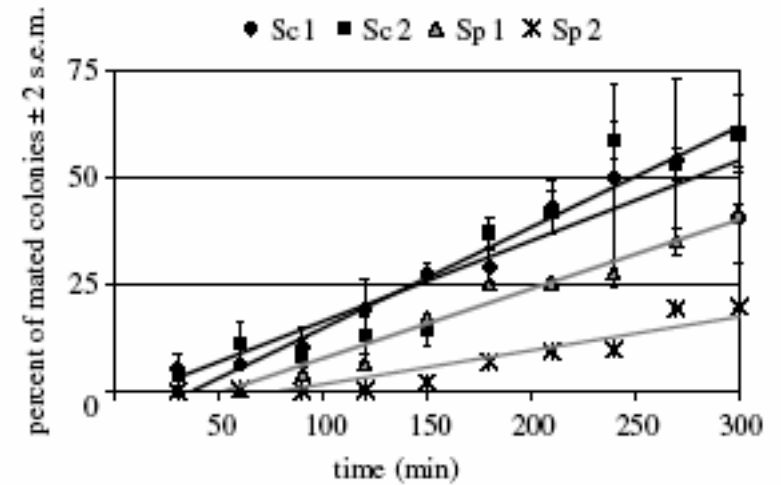
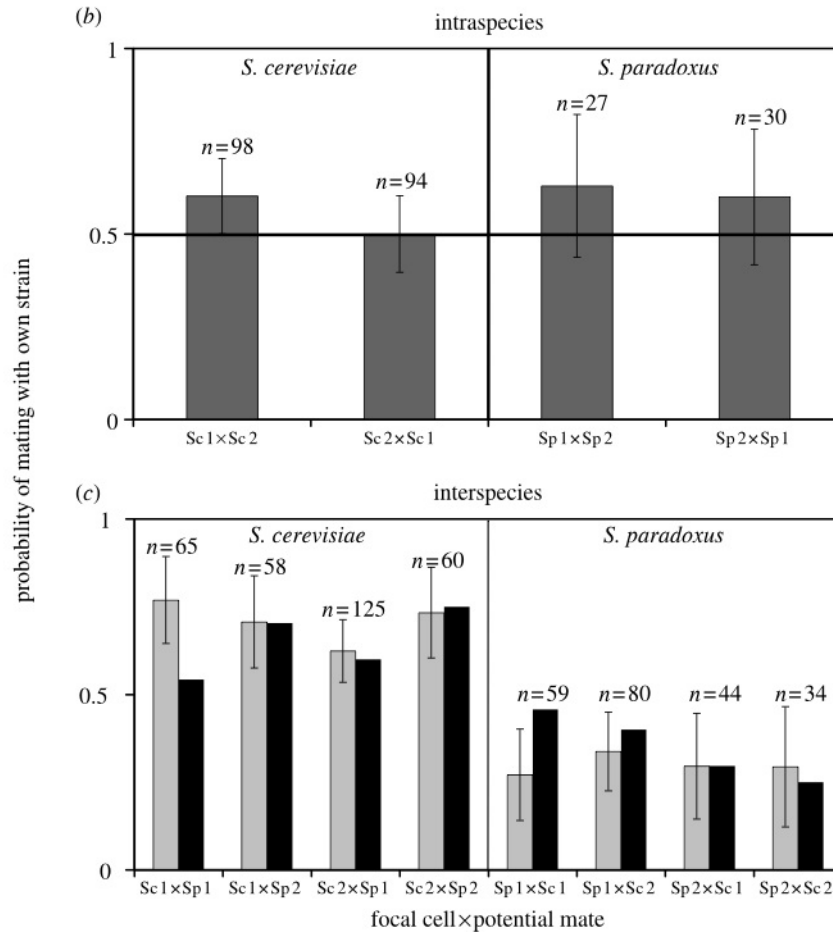
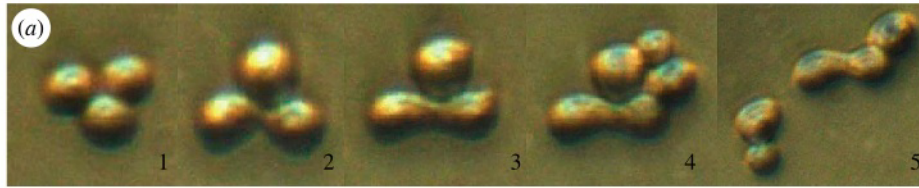
For collaboration and for generously teaching me about yeast: Gennadi Naumov and Elena Naumova.

For sage advice and encouragement: André Lachance, Western Ontario U.

For help with the work: Peter Dombrowski, Chantal Francis, Ethan Fingerman, Joe Sweeney; numerous undergraduates.

For funding: Center for Microbial Ecology, Michigan State University; University of Pennsylvania Research Foundation; National Science Foundation

Premating isolation between *cerevisiae* and *paradoxus*?



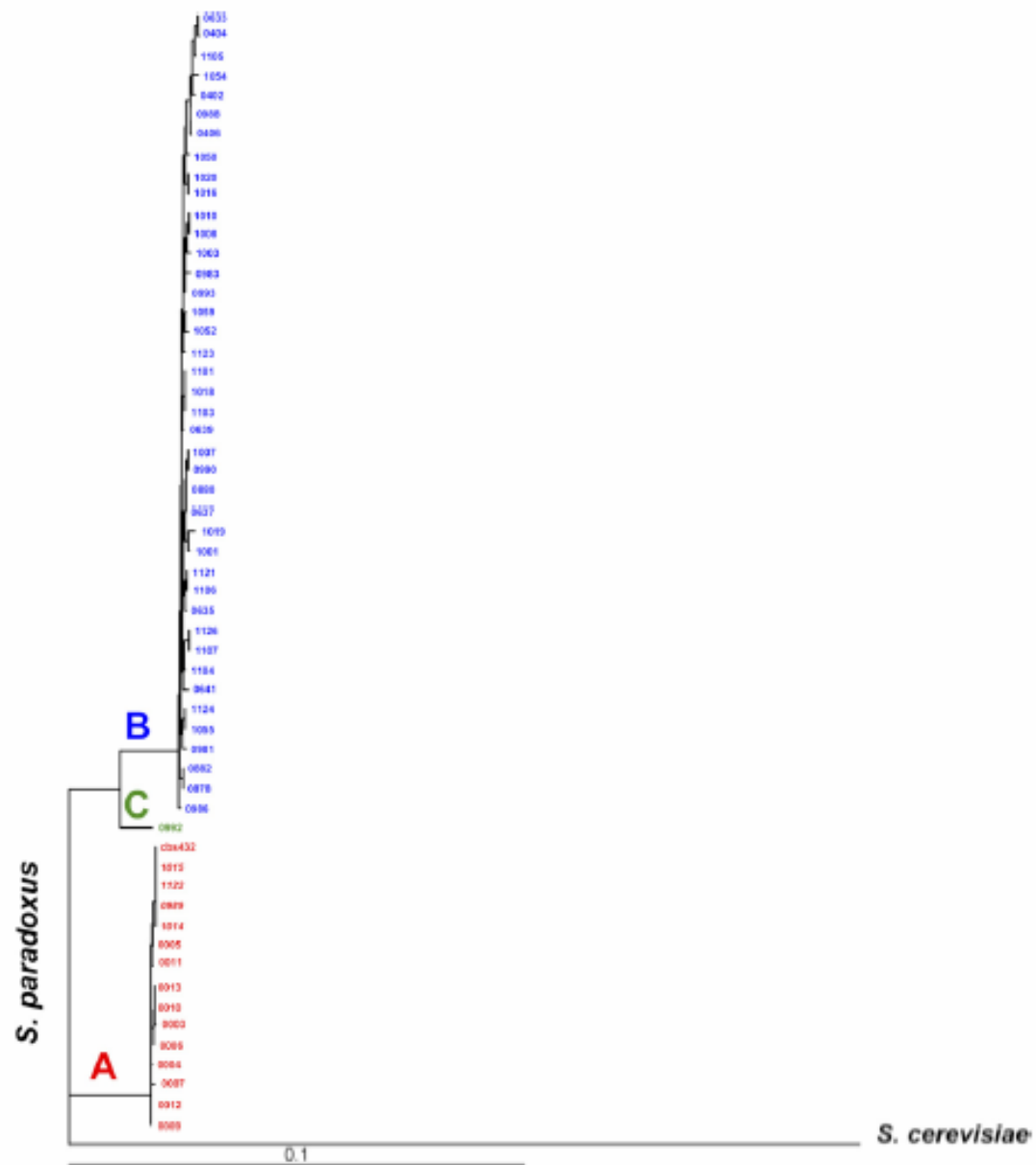


Figure S2. Complete Maximum-Likelihood Tree of the Three *S. paradoxus* Genetic Groups Rooted on *S. cerevisiae*

The tree was constructed with nine concatenated intron sequences from 57 *S. paradoxus* isolates; orthologous *S. cerevisiae* sequences were obtained from the Saccharomyces Genome Database. The major nodes separating groups A, B, and C received 100% bootstrap support (1000 replicates); the scale bar represents the number of per-site substitutions.

Different thermal growth profiles in *paradoxus* and *cerevisiae*

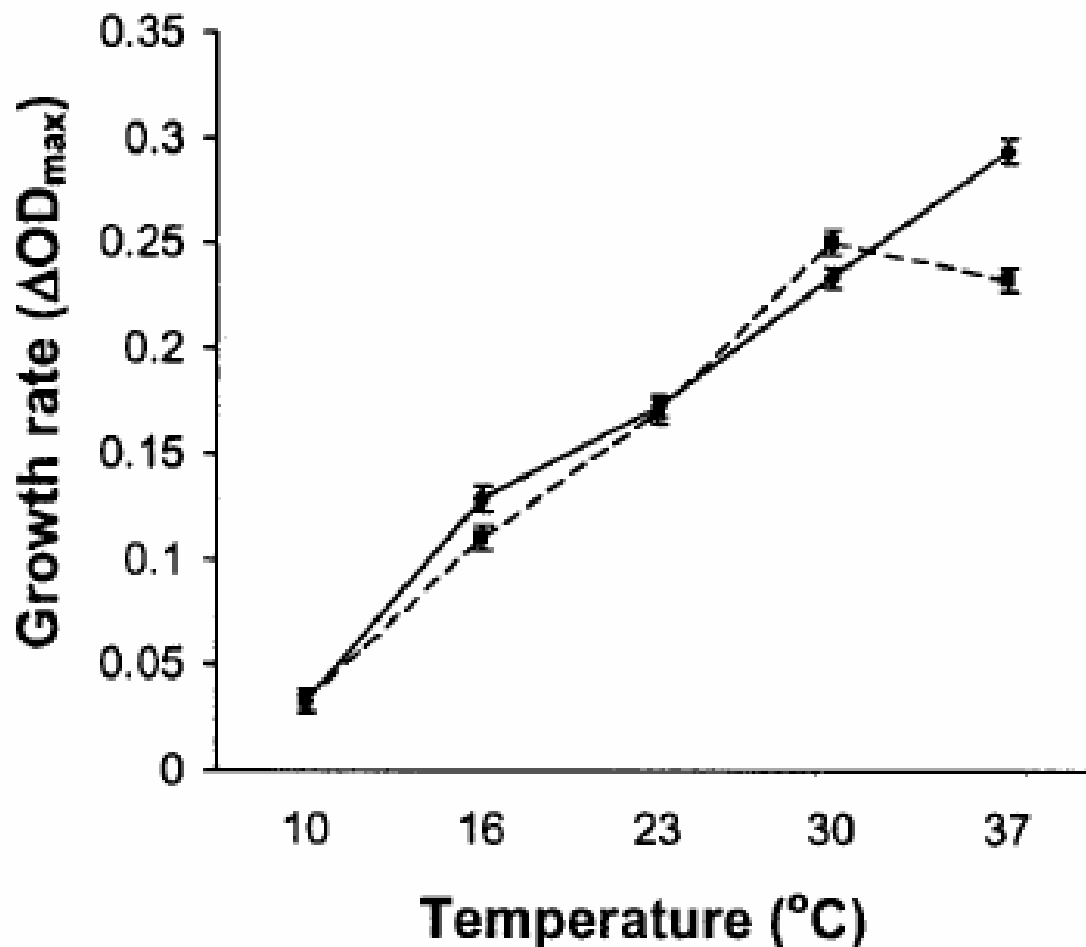


Fig. 1. Average growth rates (\pm S.E.M.) of sympatric natural *S. cerevisiae* (solid line) and *S. paradoxus* (dashed line) isolates at five temperatures. Growth rate values, shown as ΔOD_{max} , are the maximal hourly change in A_{600} observed during culture growth.