

Supplemental Information

for

Priyadarshini, Y. and Natarajan K. (2016).

Reconfiguration of Transcriptional Control of Lysine Biosynthesis in

Candida albicans Involves a Central Role for the Gcn4

Transcriptional Activator.

mSphere *In Press*

Supplementary Table S1. List of plasmids and strains

Plasmid	Relevant Description	Source
pGAL-HA	HA ₃ tagging plasmid in pESC-LEU	Sawhney and Natarajan
pGH14-2	P _{GAL1} .CaLYS141-HA ₃ , LEU2	This work
pGH142-1	P _{GAL1} .CaLYS142-HA ₃ , LEU2	This work
pGH143-9	P _{GAL1} .CaLYS143-HA ₃ , LEU2	This work
pGH144-1	P _{GAL1} .CaLYS144-HA ₃ , LEU2	This work
pYPC24	Clp10-LEU2+CaGCN4	This work
pHAH	HAH1 disruption cassette	(1)
<i>S. cerevisiae</i> strain	Relevant Genotype	Source
BY4741	<i>MATa his3Δ1 leu2Δ0 met15Δ0 ura3Δ0</i>	(2)
3973	<i>MATa his3Δ1 leu2Δ0 met15Δ0 ura3Δ0 lys14Δ::kanMX4</i>	Open Biosystems
PY1-a	[pGH14-2] in strain 3973	This work
PY9	[pGH142-1] in strain 3973	This work
PY13	[pGH143-9] in strain 3973	This work
PY17	[pGH144-1] in strain 3973	This work
249	<i>MATa his3Δ1 leu2Δ0 met15Δ0 ura3Δ0 gcn4Δ::kanMX4</i>	Open Biosystems
KNY164	<i>MATα leu2-3, 112 ura3-52 ino1 trp1::hisG</i>	(3)
KNY124	<i>MATα leu2-3, 112 ura3-52 ino1 trp1::hisG gcn4Δ-103</i>	(3)
<i>C. albicans</i> Strain	Relevant Genotype	Source
SN152	<i>arg4Δ/arg4Δ leu2Δ/leu2Δ his1Δ/his1Δ URA3/ura3Δ::imm⁴³⁴ IRO1/iro1Δ::imm⁴³⁴</i>	(4)
RPC206	<i>his1Δ/his1Δ::<pFA-HIS1> RPS1/rps1::<Clp10-C.d.ARG4> derived from SN152</i>	(5)
PC2	<i>LYS141/lys141Δ::C.d. ARG4</i> derived from SN152	This work
PC6	<i>LYS142/lys142Δ::C.d. ARG4</i> derived from SN152	This work
PC9	<i>LYS143/lys143Δ::C.d. ARG4</i> derived from SN152	This work
PC12	<i>LYS144/lys144Δ::C.d. ARG4</i> derived from SN152	This work
PC20	<i>lys141Δ::C.d.HIS1/lys141Δ::C.d.ARG4</i> derived from PC2	This work
PC27	<i>lys142Δ::C.d.HIS1/lys142Δ::C.d.ARG4</i> derived from PC6	This work
PC31	<i>lys143Δ::C.d. HIS1/lys143Δ::C.d. ARG4</i> derived from PC9	This work
PC37	<i>lys144Δ::C.d. HIS1/lys144Δ::C.d.ARG4</i> derived from PC12	This work
PC43	<i>lys2Δ::C.d. HIS1/lys2Δ::C.d.ARG4</i> derived from SN152	This work

PC205	<i>LYS141::TAP-C.m.LEU2/lys141Δ::C.d.ARG4</i> derived from PC2	This work
PC206	<i>LYS142::TAP-C.m.LEU2/lys142Δ::C.d.ARG4</i> derived from PC6	This work
PC209	<i>LYS143::TAP-C.m.LEU2/lys143Δ::C.d.ARG4</i> derived from PC9	This work
PC212	<i>LYS144::TAP-C.m.LEU2/lys144Δ::C.d.ARG4</i> derived from PC12	This work
PC222	<i>GCN4::TAP-CmLEU2/ GCN4</i> derived from SN152	This work
PC227	<i>GCN4::TAP-C.m.LEU2/ GCN4::TAP-C.d.HIS1</i> derived from PC222	This work
PC234	<i>gcn4Δ::C.d.ARG4/gcn4Δ::C.d.HIS1</i> derived from SN152 * the strain requires Arg for growth, likely due to the <i>gcn4</i> mutation	This work
PC291	<i>CaGCN4</i> (pYPC24) integrated in PC234	This work

Supplementary Table S2. List of oligonucleotides

Oligo Name	Sequence (5'-3')	Notes
ON364	5'- CGAATCAAATTAACAACCATAGGA -3'	<i>ScGAL1</i> promoter
ONC47	5'- ccggaattCTATTGCTACCACAGTCA -3'	5' cloning primer for <i>CaGCN4</i>
ONC110	5'- cggtgccacatttcacac-3'	3' Primer for sequencing <i>HIS1-TAP</i>
ONC164	5' CCGctcgagGCACTGAGCAGCGTAATC-3'	Rev Primer for HA tagging
ONC305	5'-TAC CAA GGA ACA CCA ATA TCT AGT CAC T-3'	Forward Primer for Chr1
ONC306	5'-GAT CAC TTT TGT CCT TGG CAG TC-3'	Reverse Primer for Chr1
ONC350	5'-cgggaATGTCACAATCACCATCT-3'	5' cloning primer for <i>CaLYS141</i>
ONC351	5'-tttgtcgacGTATATCAATGTATCATCATTT-3'	3' cloning primer for <i>CaLYS141</i>
ONC352	5'-cgggaATGTCTACTTCCAAGAGA-3	5' cloning primer for <i>CaLYS142</i>
ONC353	5'-TTTgtcgacGTAGGCAACATTATCGAT	3' cloning primer for <i>CaLYS142</i>
ONC354	5'-cgggaATGGCGTCAAATATTTCA-3'	5' cloning primer for <i>CaLYS143</i>
ONC355	5'-TTTgtcgacGTATGATAACAAATCCTCG-3'	3' cloning primer for <i>CaLYS143</i>
ONC356	5'-cgggaATGGGTTCATCTAGCCCAAACC-3'	5' cloning primer for <i>CaLYS144</i>
ONC357	5'-TTTgtcgacTGAAAATGGTATAATTTCCGAATCAAA-3'	3' cloning primer for <i>CaLYS144</i>
ONC362	5'-GGTGCTGATTTGAATGTTTTAAC-3'	Forward Internal primer for <i>LYS14</i>
ONC363	5'-GACCAACAACCTTTAATACTGAAAC-3'	Forward Internal primer for <i>LYS142</i>

ONC364	5'-TTGTTGTGGGCACGGAAT-3'	Reverse Internal primer for <i>LYS142</i>
ONC365	5'-CCATCAATATCATCTAACTCATCG-3'	Forward Internal primer for <i>LYS143</i>
ONC366	5'-TCGATCCCTAATTTGTTGAATG-3'	Forward Internal primer for <i>LYS143</i>
ONC367	5'-AGGATATTACTTGAGACTTGTTG-3'	Reverse Internal primer for <i>LYS143</i>
ONC368	5'-AATGATCAAGCTTTCCTAAT-3'	Forward Internal primer for <i>LYS144</i>
ONC369	5'-ACCATCTACAACATTATCACCG-3'	Forward Internal primer for <i>LYS144</i>
ONC370	5'-TTGTTCACTCCAATCGTACC-3'	Reverse Internal primer for <i>LYS144</i>
ONC378	5'-ATCAACCAACTATCATCCCAACCACTACA ACTATTCATAAA C CAA G TACTGACTTCGTCCTTCGTACGCTGCAGGTC-3'	5' Primer with 60bp homology to <i>LYS14</i>
ONC379	5'-ACTAAAACAGTACGTACAGACTTTTAAATGTTTTCCTAATTG TAGACA ACCAAATATTTTCTGATATCATCGATGAATTTCGAG-3'	3' Primer with 60bp homology to <i>LYS14</i>
ONC380	5'- GCTTTGTTTTGTTTGTGTTTCCT -3'	5' upcheck Primer for <i>LYS141</i>
ONC381	5'- TCCCTGTCGCCAGAGAAC -3'	3' downcheck Primer for <i>LYS141</i>
ONC382	5'-AAGCGGAGTTGATTTCCCTGTAGTTGTCAAATTCATATTCAC CAGCACCAAGATATCATCCTTCGTACGCTGCAGGTC-3'	5' Primer with 60bp homology <i>LYS142</i>
ONC383	5'-GATGGTAGTGTTATTTCAAATGATCCATCTTTGTCAACTTAC AAAGAATAAATACCCCTACTGATATCATCGATGAATTTCGAG-3'	3' Primer with 60 bp homology to <i>LYS142</i>
ONC384	5'- GTTGGTCTTTCGAATATTTGGG -3'	5' upcheck Primer for <i>LYS142</i>
ONC385	5'- ATGAGGTAAGAAATGAAGAGCC -3'	3' downcheck Primer for <i>LYS142</i>

ONC386	5'-CCTGAATCCAGAAAGAGGCCCATACTATTATCATATCCCCA TATCTTTATTAGTTTATCTCTTCGTACGCTGCAGGTC-3'	5'Primer with 60 bp homology to <i>LYS143</i>
ONC387	5'-TATTTAATAATACAAAATTCTAATTTTATTTAAAAGCTATA AACATATACATCT GTAGTCTGATATCATCGATGAATTCGAG-3'	3' Primer with 60 bp homology to <i>LYS143</i>
ONC388	5'- AACTAAACAAGCGAAAAAAAAAAGAG -3'	5' upcheck Primer for <i>LYS143</i>
ONC389	5'- TCATTTGGTAGAGCAAGTTATG -3'	3' downcheck Primer for <i>LYS143</i>
ONC390	5'-AGGCATCATTATTCCTTTCCATAACTCAAACCAATTACAGT AATTGAAGATAA GATCAACTTCGTACGCTGCAGGTC-3'	5'Primer with 60 bp homology to <i>LYS144</i>
ONC391	5'-ACAAAATCTATAAAACCGTGACACACAAAACCTAACTTACA CTTACAAGCAATATCTATTCTGATATCATCGATGAATTCGAG-3'	3' Primer with 60bp homology to <i>LYS144</i>
ONC392	5'- AAAATCCCCAATGACCGC-3'	5' upcheck Primer for <i>LYS144</i>
ONC393	5'- CATATGTGTGTTTGACAGTTTTTC -3'	3' downcheck Primer for <i>LYS144</i>
ONC394	5'-ATTTTATTTTCAATTTCTTTATTTGAAATAACTTTAACGACCTT CAATTAATTCACAACACTTCGTACGCTGCAGGTC-3'	5'Primer with 60bp homology to <i>LYS2</i>
ONC395	5'-TATATACTACAACCTAAGACATAAGTTCTCCTTTTCTGGTATT TTAACTTTGAAGTATTACCTGATATCATCGATGAATTCGAG-3'	3'Primer with 60bp homology to <i>LYS2</i>
ONC396	5'- CTTGTACTGTTTTGCACGAC-3'	5' upcheck Primer for <i>LYS2</i>
ONC397	5'- ACACATTGTAACATGGTAATAGC -3'	3' downcheck Primer for <i>LYS2</i>
ONC460	5'-TTTACAAACCGATGCCGAAAT-3'	5' qRT-PCR Primer for <i>LYS141</i>
ONC461	5'-TCTCTCAACACTTCAATCTCAAATTCA-3'	3' Real time Primer for <i>LYS141</i>
ONC462	5'-TGTCTATTACCAGGACCAATTTCCA-3'	5' qRT-PCR Primer for <i>LYS142</i>
ONC463	5'-TGTGGGCACGGAATCGA-3'	3' qRT-PCR Primer for <i>LYS142</i>

ONC464	5'-GCAAGCAACGGTACTGCTGAT-3'	5' qRT-PCR Primer for <i>LYS143</i>
ONC465	5'-AACGACATCCTTCTTGTGTTGAATT-3'	3' qRT-PCR Primer for <i>LYS143</i>
ONC466	5'-AACGTGGTTTTTTTGCAATTGAA-3'	5' qRT-PCR Primer for <i>LYS144</i>
ONC467	5'-AATGGTTCCGCCCAAAGG-3'	3' qRT-PCR Primer for <i>LYS144</i>
ONC468	5'-TCCGAAGCTGACGATTTACAAG-3'	5' qRT-PCR Primer for <i>LYS2</i>
ONC469	5'-TGGATTGTCCATATCCGTTTCC-3'	3' qRT-PCR Primer for <i>LYS2</i>
ONC540	5'-TGGGTTATTGAAAGATGGAGAAGTT-3'	5' qRT-PCR Primer of <i>LYS4</i>
ONC541	5'-CAATGCATCTTTTGAACCCATTC-3'	3' qRT-PCR Primer of <i>LYS4</i>
ONC548	5'-CTGGCGGTAAAATCGTATCATTCC-3'	5' qRT-PCR Primer of <i>LYS9</i>
ONC549	5'-AATAAAACCCCTCTTGAACTCAA-3'	3' qRT-PCR Primer <i>LYS9</i>
ONC561	5'-AGTATTATGATCTTCATATTAAGCATGAAAATGGTCAAATT GACGGAATTCATATCGATAATGTTGCCTACGGATCCATGGAAAAGAGA- 3'	5' Primer for c-term TAP tagging Primers for <i>CaLYS142</i>
ONC562	5'-ATATATCCAGATGGTAGTGTATTTCAAATGATCCATCTTTG TCAACTTACAAAGAATAAATACCCCTAGTGTGATGGATATCTGCA3'	3' Primer for C-term TAP tagging Primers for <i>CaLYS142</i>
ONC563	5'-TGGAATAAAAGAGATAATATCGAAGAAAGCGATAACGAT GACAACGAGAACGAGGATTTGTTATCATAACGGATCCATGGAAAAGAGA- 3'	5' Primer for c-term TAP tagging Primers for <i>CaLYS143</i>
ONC564	5'-TTTTAAGAATATTTAATAATACAAAATTCTAATTTTATTTTA AAAGCTATAAACATATAACATCTGTAGTGTGTGATGGATATCTGCA	3' Primer for c-term TAP tagging Primers for <i>CaLYS143</i>
ONC565	5'-TCTATACAAATCTTACTAAAACATTGGAAAGGGACAATTGA TATTGAAAATGATGATACATTGATATACGGATCCATGGAAAAGAGA-3"	5' Primer for c-term TAP tagging Primers for <i>CaLYS14</i>
ONC566	5'-AAATAAAGAACTAAAACAGTACGTACAGACTTTTAAATGTT TTCTAATTGTAGACAACCAAATATTTTGTGTGATGGATATCTGCA-3'	3' Primer for c-term TAP tagging Primers for <i>CaLYS14</i>
ONC567	5'-TTCATTGACGGTAAAGATGTAGATGATGACGTTGATTTAGC ATTTGATTCGGAAATTATACCATTTTCAGGATCCATGGAAAAGAGA-3'	5' Primer for c-term TAP tagging Primers for <i>CaLYS144</i>

ONC568	5'-AATAACAAAACAAAATCTATAAAACCGTGACACACAAAAC TAAACTTACTTACAAGCAATATCTATTGTGTGATGGATATCTGCA-3'	3' Primer for c-term TAP tagging Primers for <i>CaLYS144</i>
ONC620	5'-GCGTTTCGTGGAGTCATTGAA	5' ChIP Primer for <i>LYS2</i>
ONC621	5'-GCACGACTTCCAATTATCAAGGA	3' ChIP Primer for <i>LYS2</i>
ONC622	5'-CTACCAAGTGACTCATTAGAGGCG	5' ChIP Primer for <i>LYS2</i>
ONC623	5'-TGTGACTAATCAAAGAAAGCGGC	3' ChIP Primer for <i>LYS2</i>
ONC624	5'-TCACTTGTTCTACTCTTTCTTCTTTTTTC	5' ChIP Primer for <i>LYS9</i>
ONC625	5'-ACCCCTTCCACCAACTCTCTC	3' ChIP Primer for <i>LYS9</i>
ONC626	5'-GGGAAGTGATAAGATTGCCGC	5' ChIP Primer for <i>LYS9</i>
ONC627	5'-CGTGTATTTATGGTAGATAATGAGTTGACTC	3' ChIP Primer for <i>LYS9</i>
ONC628	5'-GCCGCATATGCTATTACATTG	5' ChIP Primer for <i>LYS9</i>
ONC629	5'-CAATAAGCTCTAATTCAATGATGACATG	3' ChIP Primer for <i>LYS9</i>
ONC634	5'-GAAAAGCAAGCTTTACAAGATCAAGTTGAAAGATTACAAG AATTGTTAAGAGTTAATGGTATTCAATTTGGATCCATGGAAAAGAGA-3'	5' Primer for TAP-tagging of <i>GCN4</i>
ONC635	5'-AACATAATAATTTCTAAATTTTCTTTTTTAAAAAATA ACGAGAGGTATATATAGTAGTAACTTTGTGTGATGGATATCTGCA	3' Primer for TAP-tagging of <i>GCN4</i>
ONC657	5'- TTATTATTTACTGTTGCTATTTAAATTAATTACATTA CATTAATTAGCTTTGTTACCCTTCGTACGCTGCAGGTC -3'	5' Primer with 60bp homology to <i>GCN4</i>
ONC658	5'-TAACGAGAGGTATATATAGTAGTAACTTTTAAATTTGAAT ACCATTA ACTCTTAACAATCTGATATCATCGATGAATTCGAG-3'	3' Primer with 60bp homology to <i>GCN4</i>

ONC659	5'-ACTACTTAGAAAGTCCTG -3'	5' Upcheck Primer of Ca <i>GCN4</i>
ONC660	5'- ATACTATCTGGACGAAGA -3'	3' Downcheck Primer of Ca <i>GCN4</i>
ONC1011	5'- TTT GTC GAC GAC TAT CCT ACA CTC A -3'	Forward primer for cloning <i>GCN4</i>
ONC1012	5'- ATC GAT GTC TAC CAT CAA TTA AAC T -3'	Reverse primer for cloning <i>GCN4</i>

Supplementary Table S3. Location of single nucleotide differences in the *LYS14* coding sequences and their alleles in the *C. albicans* genome sequence assembly and the corresponding sequence in the cloned *LYS14* genes.

Hap A Chromosome	Hap A Upstream Coord	Hap A Downstream Coord	Hap B Chromosome	Hap B Upstream Coord	Hap B Downstream Coord	Change Type	Hap A Seq	Hap B Seq	Affected Features	Amino acid Change: Y/N; residue	Nucleotide change	Amino acid change
Ca22chr6A_C_albicans_SC5314	567324	567326	Ca22chr6B_C_albicans_SC5314	567295	567297	1 nt subs	CAG	CGG	C6_02750C/LYS14		CGG	
Ca22chr1A_C_albicans_SC5314	2020319	2020321	Ca22chr1B_C_albicans_SC5314	2020354	2020356	1 nt subs	GCT	GTT	C1_09200W/LYS142		GCT	
Ca22chr1A_C_albicans_SC5314	2020373	2020375	Ca22chr1B_C_albicans_SC5314	2020408	2020410	1 nt subs	CTA	CCA	C1_09200W/LYS142		CTA	
Ca22chr1A_C_albicans_SC5314	2020896	2020898	Ca22chr1B_C_albicans_SC5314	2020931	2020933	1 nt subs	GCC	GTC	C1_09200W/LYS142	P233S	GTC	P233S
Ca22chr1A_C_albicans_SC5314	2021018	2021020	Ca22chr1B_C_albicans_SC5314	2021053	2021055	1 nt subs	GCG	GGG	C1_09200W/LYS142		GCG	
Ca22chr1A_C_albicans_SC5314	2021123	2021125	Ca22chr1B_C_albicans_SC5314	2021158	2021160	1 nt subs	GGG	GTG	C1_09200W/LYS142		GGG	
Ca22chr1A_C_albicans_SC5314	2021132	2021134	Ca22chr1B_C_albicans_SC5314	2021167	2021169	1 nt subs	TTT	TAT	C1_09200W/LYS142		TTT	
Ca22chr1A_C_albicans_SC5314	2022097	2022099	Ca22chr1B_C_albicans_SC5314	2022132	2022134	1 nt subs	GGG	GAG	C1_09200W/LYS142	G633E	GGG	
Ca22chr1A_C_albicans_SC5314	2015059	2015061	Ca22chr1B_C_albicans_SC5314	2015094	2015096	1 nt subs	AAG	AGG	C1_09180W/LYS143		AAG	
Ca22chr1A_C_albicans_SC5314	2015148	2015150	Ca22chr1B_C_albicans_SC5314	2015183	2015185	1 nt subs	ATA	ACA	C1_09180W/LYS143	I109T	ATA	
Ca22chr1A_C_albicans_SC5314	2015215	2015217	Ca22chr1B_C_albicans_SC5314	2015250	2015252	1 nt subs	CCC	CTC	C1_09180W/LYS143		CCC	
Ca22chr1A_C_albicans_SC5314	2015257	2015259	Ca22chr1B_C_albicans_SC5314	2015292	2015294	1 nt subs	AAC	AGC	C1_09180W/LYS143		AAC	
Ca22chr1A_C_albicans_SC5314	2015272	2015274	Ca22chr1B_C_albicans_SC5314	2015307	2015309	1 nt subs	CGC	CTC	C1_09180W/LYS143		CGC	
Ca22chr1A_C_albicans_SC5314	2015321	2015323	Ca22chr1B_C_albicans_SC5314	2015356	2015358	1 nt subs	AGT	ACT	C1_09180W/LYS143		AGT	
Ca22chr1A_C_albicans_SC5314	2015473	2015480	Ca22chr1B_C_albicans_SC5314	2015508	2015515	6 nt subs	GGCCACA C	GCCCACC C	C1_09180W/LYS143	H219P	GGCCACAC	
Ca22chr1A_C_albicans_SC5314	2015621	2015623	Ca22chr1B_C_albicans_SC5314	2015656	2015658	1 nt subs	ATT	ACT	C1_09180W/LYS143		ATT	
Ca22chr1A_C_albicans_SC5314	2015641	2015643	Ca22chr1B_C_albicans_SC5314	2015676	2015678	1 nt subs	CGG	CAG	C1_09180W/LYS143		CGG	
Ca22chr1A_C_albicans_SC5314	2015671	2015673	Ca22chr1B_C_albicans_SC5314	2015706	2015708	1 nt subs	CCG	CTG	C1_09180W/LYS143		CCG	
Ca22chr1A_C_albicans_SC5314	2015694	2015696	Ca22chr1B_C_albicans_SC5314	2015729	2015731	1 nt subs	GCT	GTT	C1_09180W/LYS143		GCT	

Ca22chr1A_C_albicans_SC5314	2015704	2015706	Ca22chr1B_C_albicans_SC5314	2015739	2015741	1 nt subs	GCA	GTA	C1_09180W/LYS143		GCA	
Ca22chr1A_C_albicans_SC5314	2015716	2015718	Ca22chr1B_C_albicans_SC5314	2015751	2015753	1 nt subs	GCA	GTA	C1_09180W/LYS143		GCA	
Ca22chr1A_C_albicans_SC5314	2015743	2015745	Ca22chr1B_C_albicans_SC5314	2015778	2015780	1 nt subs	ATG	ACG	C1_09180W/LYS143		ATG	
Ca22chr1A_C_albicans_SC5314	2016601	2016603	Ca22chr1B_C_albicans_SC5314	2016636	2016638	1 nt subs	CCC	CGC	C1_09180W/LYS143		CGC	
Ca22chr1A_C_albicans_SC5314	2016764	2016766	Ca22chr1B_C_albicans_SC5314	2016799	2016801	1 nt subs	ATT	ACT	C1_09180W/LYS143		ACT	
Ca22chr1A_C_albicans_SC5314	2016838	2016840	Ca22chr1B_C_albicans_SC5314	2016873	2016875	1 nt subs	TCG	TTG	C1_09180W/LYS143		TTG	
Ca22chr1A_C_albicans_SC5314	2016865	2016867	Ca22chr1B_C_albicans_SC5314	2016900	2016902	1 nt subs	TCC	TTC	C1_09180W/LYS143		TTC	
Ca22chr1A_C_albicans_SC5314	2016930	2016935	Ca22chr1B_C_albicans_SC5314	2016965	2016970	4 nt subs	GCTATA	GTTACA	C1_09180W/LYS143	I704T	GTTACA	I704T
Ca22chr1A_C_albicans_SC5314	2017073	2017075	Ca22chr1B_C_albicans_SC5314	2017108	2017110	1 nt subs	TCG	TGG	C1_09180W/LYS143	R751G	TGG	R751G
Ca22chr1A_C_albicans_SC5314	2017168	2017170	Ca22chr1B_C_albicans_SC5314	2017203	2017205	1 nt subs	TGC	TAC	C1_09180W/LYS143		TAC	
Ca22chr3A_C_albicans_SC5314	140495	140510	Ca22chr3B_C_albicans_SC5314	140494	140506	3 nt indel	TACCACC ACCACCA CA	TACCACC ACCACA	C3_00750W/LYS144	T22_	TACCACCA CCACCA	
Ca22chr3A_C_albicans_SC5314	141210	141212	Ca22chr3B_C_albicans_SC5314	141206	141208	1 nt subs	GGA	GAA	C3_00750W/LYS144	G260E	GGA	
Ca22chr3A_C_albicans_SC5314	141649	141651	Ca22chr3B_C_albicans_SC5314	141645	141647	1 nt subs	TTC	TCC	C3_00750W/LYS144		TTC	
Ca22chr3A_C_albicans_SC5314	141764	141766	Ca22chr3B_C_albicans_SC5314	141760	141762	1 nt subs	TCT	TTT	C3_00750W/LYS144		TCT	
Ca22chr3A_C_albicans_SC5314	141805	141807	Ca22chr3B_C_albicans_SC5314	141801	141803	1 nt subs	ATA	ACA	C3_00750W/LYS144		ATA	
Ca22chr3A_C_albicans_SC5314	142285	142287	Ca22chr3B_C_albicans_SC5314	142281	142283	1 nt subs	GCG	GTG	C3_00750W/LYS144		GCG	
Ca22chr3A_C_albicans_SC5314	142711	142715	Ca22chr3B_C_albicans_SC5314	142707	142711	3 nt subs	ACACG	ATAAG	C3_00750W/LYS144	T761K	ACACG	
Ca22chr3A_C_albicans_SC5314	142838	142840	Ca22chr3B_C_albicans_SC5314	142834	142836	1 nt subs	TAG	TGG	C3_00750W/LYS144	S803G	TAG	
Ca22chr3A_C_albicans_SC5314	142969	142971	Ca22chr3B_C_albicans_SC5314	142965	142967	1 nt subs	ATG	AGG	C3_00750W/LYS144		ATG	
Ca22chr3A_C_albicans_SC5314	143018	143020	Ca22chr3B_C_albicans_SC5314	143014	143016	1 nt subs	GAA	GCA	C3_00750W/LYS144		nt subs after stop codon	

Supplementary Table S4. Location of CTG codons in *C. albicans* *LYS14*-like Genes

CaLys14-like sequence	No. of CTG codons/ Leu-to-Ser in <i>S.cerevisiae</i>	Amino acid Position of CTG → Ser in the translated protein	Zinc Finger domain
Lys141	1	22	32~82
Lys142	5	86, 168, 347, 399, 610	5~55
Lys143	6	68, 86, 130, 184, 781, 835	20~70
Lys144	8	95, 123, 286, 304, 360, 554, 613, 793	36~86

Supplementary Figure Legends

FIG S1. *S. cerevisiae* lysine biosynthetic pathway genes are induced under amino acid starvation conditions. Summary of genome-wide microarray data of *S. cerevisiae* *LYS* genes under His starvation (+/-3AT) (6) or under Ile-Val starvation (+/- SM) conditions (7).

FIG S2. *CaLYS14*-like genes do not complement the lysine auxotrophy of *S. cerevisiae* *lys14Δ* strain. *S. cerevisiae* *lys14Δ* strain was transformed with plasmids containing *P_{GALI}*-driven *CaLYS141*, *CaLYS142*, *CaLYS143* or *CaLYS144* and the transformants as well as the control strains wild-type BY4741 and the parental *lys14Δ* (strain 3973) were grown overnight in SC medium at 30°C. Saturated cultures were diluted and spotted on synthetic complete media containing glucose or galactose plus raffinose as carbon sources and with or without lysine supplementation.

FIG S3. Deletion of *CaLYS14*-like genes does not lead to lysine auxotrophy in *C. albicans*. *C. albicans* strain RPY206 (WT) and the different mutant strains *lys141Δ/Δ*, *lys142Δ/Δ*, *lys143Δ/Δ*, *lys144Δ/Δ*, and *lys2Δ/Δ* as control, were grown overnight in YPD, and 10-fold serial dilutions were spotted on SC-Lys, SC-Lys+0.1mM Hyl, or on control plate SC+Lys+0.1mM Hyl, and grown for 2 days and image acquired.

FIG S4. Lysine starvation does not lead to upregulation of *C. albicans* *LYS14* gene expression. The wild-type strain RPC206 was grown in SC+Lys (lysine-replete), SC-Lys (lysine-deficient) or SC-Lys+Hyl (lysine-starved) media, extracted total RNA and the mRNA levels of *LYS141*, *LYS142*, *LYS143* and *LYS144* were quantified using qRT-PCR. The *scR1* transcript was used as normalization control, and relative expression calculated with respect to SC+lys, and data from 2-3 independent experiments were plotted. The error bars indicate SEM.

FIG S5. CaLys14-like protein levels are not altered by lysine starvation conditions. *CaLYS14* genes were TAP-tagged at their carboxyl-termini and the tagged strains PC205, 206, 209 and 212 expressing Lys141-TAP, Lys142-TAP, Lys143-TAP and Lys144-TAP respectively, were grown in SC+Lys, SC-Lys and SC-Lys +Hyl to OD₆₀₀ ~2.0, and harvested. Whole cell extracts were prepared and 2-fold serial dilutions, starting from 50µg, were analyzed by Western blot. The TAP-tagged Lys14 expression was probed with PAP antibody (1:6000; Open Biosystems) and anti-Glucose 6-phosphate dehydrogenase antibody (1:1000; Sigma) as loading control.

FIG S6. Comparison of the *S. cerevisiae* and *C. albicans* *LYS* biosynthetic gene promoters. Schematic diagram comparing the presence or absence of the binding sites for Lys14 and Gcn4 on the promoters of the lysine biosynthetic pathway genes in *S. cerevisiae* and *C. albicans* based on their analyses at the yTFaSCo database.

FIG S7. Comparison of the *LYS* biosynthetic gene promoters in *Candida* clade genomes. Schematic diagram comparing the presence or absence of the binding sites for Lys14 and Gcn4 on the promoters of the lysine biosynthetic pathway genes in *S. cerevisiae*, *C. albicans* and other *Candida* clade genomes *L. elongisporus*, *C. tropicalis*, *C. lusitaniae*, and *C. guilliermondii* based on their analyses at the yTFaSCo database.

Supplementary References

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	WT +/- SM	WT +/- 3AT
	log2fc	log2fc
<i>LYS20</i>	2.8	3.0
<i>LYS21</i>	1.5	2.4
<i>LYS4</i>	1.8	1.6
<i>LYS12</i>	2.1	n.d.
<i>LYS14</i>	1.9	2.7
<i>YER152C</i>	no change	n.d.
<i>ARO8</i>	1.1	2.5
<i>ARO9</i>	4.1	5.4
<i>LYS2</i>	2.8	3.4
<i>LYS5</i>	2.2	2.7
<i>LYS9</i>	2.9	2.5
<i>LYS1</i>	3.8	4.5











