Supplementary File:

Genes, Transcription Factors, Protein Kinases, Intracellular Proteins, and Other Abbreviations as Used in the Article

Aap – amino acid permease

Aat – amino acid transporter

Aca - adenylyl cyclase-associated protein

Acc1 - acetyl-CoA decarboxylase

Acl1 - ATP-citrate lyase

Aco1 - aconitase

Acs1 – acetyl-CoA synthetase for histone acetylation

Acs2/Kbc1 - 2-ketobutyryl-CoA synthetase

Ada2 - histone acetyltransferase complex SAGA/ADA

Ady2 – acetate transporter

Afr1 – ATP-binding cassette transporter

Ags – α -1,3-glucan synthase

Agt1 – α -glucoside transporter or maltose permease

Ald1 - aldehyde dehydrogenase

Ald5 - aldehyde dehydrogenase

AMPK – 5' adenosine monophosphate-activated protein kinase

Amt1/Mep1 – ammonium permease/transporter

Aox1 - alternative oxidase

Apc – amino acid-polyamine-organocation secondary carrier protein (APC) superfamily (including the amino acid-polyamine-organocation (APC), the amino acid/auxin permease (AAAP), the alanine or glycine:cation symporter (AGCS), the cation-chloride cotransporter (CCC), the hydroxy/aromatic amino acid permease (HAAAP), the betaine/carnitine/choline transporter (BCCT), the solute:sodium symporter (SSS), the neurotransmitter:sodium symporter (NSS), the nucleobase:cation symporter-1 (NCS1), the nucleobase:cation symporter-2 or nucleobase:ascorbate transporter (NCS2), the sulfate permease (SulP) as family members)

Apn1/2 - DNA-apurinic/apyrimidinic site lyase (endonuclease) 1 and 2 function in the base excision repair pathway

Aro80 - transcription activator of genes involved in the tolerance and metabolism of aromatic amino acids

Atf1 – Cre-binding basic leucine-zipper (bZIP) domain activating transcription factor regulated by Hog1

Atg - serine/threonine-protein kinase involved in autophagy

Atg26 - UDP-glucose:sterol glucosyltransferase in the biosynthesis of sterol glucoside membrane lipids

Atg4 - C54 peptidase family protein

Atg8 – autophagy-related protein (a ubiquitin-like protein associated with phosphatidylethanolamine) involved in autophagosome

Atm1 – mitochondrial inner membrane ABC transporter/iron-sulfur clusters transporter Atm1p

Atp1 – α -subunit F₀F₁-type ATP synthase

Bck1 – bypass of *c* kinase – an activator of mitogen-activated protein kinase

Bwc - Basidiomycete white collar

Cac1 - Cryptococcus adenylyl cyclase

Cam1 - calmodulin

Can – carbonic anhydrase CO₂-sensor

Cbk – cell wall biosynthesis serine/threonine kinase (a member of RAM signalling network protein kinase belonging to the AGC kinase family)

Cbp – ubiquinol-cytochrome c chaperone

Ccc1/2 – copper transporter, a P-type ATPase

Cda – chitin deacetylase

Cdc – cell division cycle transcription factors

Cdc10 – septin aspD

Cdc11 – septin aspA

Cdc12 – septin aspC

Cdc14 – phosphoprotein phosphatase

Cdc16/23 – anaphase-promoting complex subunits protein

Cdc24 – a GEF protein mediating a conserved signalling event for thermotolerance, polarised growth, and pathogenesis via Ras1, Cdc24, Cdc42, and Pak kinase Ste20

Cdc31 - EF-hand Ca2+-binding protein

Cdc36 - CCR4-NOT transcription complex

Cdc4 - myosin regulatory light chain

Cdc40 – cell division cycle 40

Cdc420 – a paralogue of Cdc42 gene

Cdc43 – encoding β-subunit of geranylgeranyltransferase-1 (Ggtase-1), catalysing the attachment of hydrophobic isoprenoid moieties onto C-terminal CaaX motifs (cysteine, 2 aliphatic amino acids, and variable amino acid)

Cdk - cyclin-dependent kinase

Cdk8 - cyclin-dependent protein kinase c, and a suppressor of Snf1; a homologue of yeast Ssn3

Chs - chitin synthase

Cin1 – cryptococcal intersectin 1 (human endocytic ITSN1 homologue)

Cir1/2 – Cryptococcus iron regulator electron transfer flavoprotein-ubiquinone oxidoreductase homologue of *S. cerevisiae*

Clr6 – cryptic loci regulator encoding putative class I histone deacetylase that removes an acetyl group from ε -N-acetyllysine amino acid residue on a histone, thereby enhancing tighter wrapping of histones on DNA to prevent access to transcription factors and DNA expression. Clr6p is a transcriptional epigenetic repressor of the Mat2 – Mat3 region and centromere for precise heterochromatin packaging during chromosomal segregation in fission yeast

Cna/Cnb - temperature- and calcium-induced calcineurin, inhibited by FK506

Coq10 – ubiquinone-binding protein

Coq11 – a putative protein involved in FMN-dependent decarboxylation in CoQ biosynthesis

Cpk1 – putative MAP kinase Ubc3

Cpp – Cryptococcus purine/cytosine permease, e.g., Cpp2 homologues of S. cerevisiae Fcy2 and Fcy21

Cpr – constitutive pheromone receptor, e.g., *Cpra* gene is a *MATa* strain-specific gene encoding putative seven-transmembrane domain pheromone receptors with diverse functions – mating, virulence, and capsule

Cps1 – hyaluronic acid synthase/cellubiuronic synthase essential in transcellular transmigration

Cps1/10 - vacuolar carboxypeptidase yscS

CPY - carboxypeptidase Y

Cqs1/2 (Qsp1/2) - Cryptococcus quorum sensing-like molecule/quorum sensing-like protein

Crg – a regulator of G-protein signalling

CRIB – Cdc42/Rac interaction binding domain, e.g., Ste20p, which can interact via CRIB with Cdc42p, activate Rac1p and Dch2p

Crk1 - Cdk-related kinase 1

Crz1/Sp1 – cell wall transcriptional regulator known as calcineurin-responsive zinc finger 1

CSR – common stress response/regulatory genes

Csr1, Csr2, Csr3 – chitin synthase regulators

Cta11 - catalase 1

Cta12/Cat3 – catalase, homologue of *S. cerevisiae*

Ctr1 – high-affinity copper transporter

Cwc1/2 - Cryptococcus white collar photoregulator, a homologue of N. Crassa white collar 1 and 2 (Wc-1/2)

Cwc24 – pre-mRNA-splicing factor homologue of *S. pombe* Cwf24p

CWI - cell wall integrity

Ddc1 - DNA damage checkpoint protein associated with Mec1

Ena1/5/50/51/52 – Na⁺ or K⁺ P-type membrane ATPase cation transporter

Erg – ergosterol regulatory gene

Erg1 – squalene epoxidase/monooxygenase

Erg10 – acetyl-CoA acetyltransferase

Erg11 – *Cyt P*₄₅₀ lanosterol 14α -demethylase

Erg110 – Δ^{14} -sterol demethylase activity with heme-binding pocket

Erg13 – 3-hydroxy-3-methylglutaryl-CoA (HMG-CoA) synthase in mevalonate biosynthesis

Erg2 – Δ ⁸-sterol isomerase

Erg20 – polyprenyl synthetase

Erg24 – Δ^{14} -sterol reductase

Erg25 – Δ^4 -methylsterol oxidase (sterol desaturase)

Erg26 – Δ^3 -sterol dehydrogenase/3- β -hydroxysteroid dehydrogenase, a homologue of *S. cerevisiae*

Erg27 – 3-keto sterol reductase

Erg3 – Δ ⁵-sterol desaturase

Erg4 – $\Delta^{24/28}$ -sterol reductase

Erg5 – Δ^{22} -sterol desaturase

Erg50 – Δ^{22} -sterol desaturase with cyt P_{450}

Erg6 – Δ^{24} -sterol methyltransferase

Erg7 – lanosterol synthase

Erg8 – phosphomevalonate kinase

Erk – extracellular signal-regulated kinase MAPK with T-E-Y motif (Cpk1 and Mpk1)

ESR – environmental stress response/regulatory groups of genes/ER-stress response genes

Fet – iron transporter

Fet3/5 – ferroxidase/multicopper oxidase or iron transporter

Fhb1 – flavohemoglobin denitrosylase (nitric oxide dioxygenase)

Fre2/7/20 – cell surface iron reductase/ferric-chelate reductase

Ftr/Fth – high-affinity ferrous iron permease/transporter (Cft gene in C. neoformans)

Fzc31 – fungal-specific Zn₂Cys₆ DNA binding protein regarded as a core virulence transcription factor for thermotolerance, antioxidation, and mating but inhibit melanin formation

Gat – transcriptional activator capable of binding -G-A-T-A- sequence within the DNA to regulate other genes. Popular GATAs are zinc-finger protein families, such as Gat1, Gat2, and Gat3

Gcs1 – glucosylceramide synthase 1, which synthesises the membrane sphingolipid GlcCer from the C9-methyl ceramide

Gpa – G-protein α subunit

Gre2 – homologue of mammalian 3- β -hydroxysteroid dehydrogenase (gene de respuesta an estrés –stress-responsive gene)

Grx – glutathione-dependent oxidoreductase (e.g., Grx4 and Grx5 monothiol glutaredoxin in *C. neoformans* but Grx3 and Grx4 in *S. cerevisiae*)

GSR - genotoxic stress response/regulatory groups of genes

Hap1 - haem-binding transcription factor

HMSR/HMR - heavy-metal stress response/regulatory groups of genes/heavy-metal resistance

Hog1 – a *p38* family of MAPK with a T-G-Y motif

Hsp10 - mitochondrial matrix co-chaperonin ATPase inhibitor

Hsp104 – heat shock protein 104

Hsp12 – plasma membrane heat-shock protein for membrane organisation during stress conditions, homologue to Hsp122. While Hsp12 appears constitutive, Hsp122 is slightly induced in *C. neoformans*. Under the cAMP and Hog1 regulations, both chaperons are involved in polyene antifungal susceptibility

Hsp31 – DJ-1/ThiJ/Pfp1 superfamily, yeast stress-inducible homodimeric heat shock protein chaperone with methylglyoxalase activity

Hsp60 – a tetradecameric mitochondrial chaperonin

Hsp70 - multifunctional ATP-dependent heat-shock molecular chaperon expressed against stress

Hsp78 – heat shock protein chaperone

Hsp90 – ATP-dependent molecular chaperone activator of protein turnover during stress

HTG - high-temperature growth

Hxl1 - Hac1, Xbp1-Like gene 1 encoding UPR transcriptional factor

Icl1-isocitrate lyase catabolising isocitrate into succinate and glyoxylate

Idh1/Idp1/Lys1 – isocitrate/isopropylmalate dehydrogenase

Idi1 – isopentenyl diphosphate ¹-isomerase

Ilv1 – threonine deaminase in isoleucine biosynthesis

Ilv2 – acetolactate synthase for isoleucine and valine biosynthesis

Ilv3 – dihydroxyacid dehydratase in the branched-chain amino acid biosynthesis

Ilv5 – acetohydroxyacid reductoisomerase

Ipc1 - inositolphosphoceramide synthase

Ire1 – ER-resident transmembrane kinase/RNase

Jac1 – J-type **a**ccessory **c**haperone

Kcs1 – protein **k**inase *c* **s**uppressor

Kic1 – serine/threonine-protein kinase related to Pak/Ste20 protein family

Kre – β -glucan synthase

Leu1 – 2-isopropylmalate synthase

Leu2 - 3-isopropylmalate dehydrogenase - an Fe-S cluster protein required for leucine biosynthesis

Leu3 – a repressor that, together with Gcn4, regulates amino acid biosynthesis

Leu9 – α -isopropylmalate synthase II (2-isopropylmalate synthase) in leucine biosynthesis

Liv3 – a Wor1-motif DNA binding protein (orthologue of *C. albicans* Wor1)

LPL - lysophospholipase

LPTA – lysophospholipase/transacylase

Lys10 – dihydrodipicolinate synthase in lysine biosynthesis

Lys2 – aminoadipate-semialdehyde dehydrogenase

Lys2 – α -aminoadipate reductase in lysine biosynthesis

Lys21 – homocitrate synthase isozyme in lysine biosynthesis

Lys4/Lys40/Lys41/Lys42 - homoaconitase, converting homocitrate to homoisocitrate in lysine biosynthesis

Mal – hexose transporter

Man1 – phosphomannose isomerase

MAPK - mitogen-activated protein kinases such as Mpk1, a homologue of S. cerevisiae Slt2 or human Erk5

May1 - major aspartyl peptidase 1

Mbs1 – APSES-like transcription factor (Mbp1- and orthologue of Swi4-like DNA binding protein 1)

Mbs2 – orthologue to Swi6-like DNA binding protein regulating the cell cycle. It can form a complex, Mbs1/Mbs2- or Swi4/Swi6-dependent cell cycle box-binding factor

Mdr1 - multidrug resistance protein 1

Met13 - methylenetetrahydrofolate reductase

Met14 – adenylylsulfate kinase

Met17 - cysteine synthase

Met2 - homoserine transacetylase

Met3 - sulfate adenylyltransferase/ATP sulfurylase

Met30 - sulfur metabolite repression control protein and a negative regulator of sulfur amino acids genes expression for biosynthesis

Met5 - nitrite and sulphite reductase with a 4Fe-4S binding domain

Met5/10 – sulfite reductase α -subunit converting sulfite into sulfide

 $\boldsymbol{Met7-} folylpolyglutamate\ synthase$

Mp88 – T cell-stimulating mannoprotein

Mrs4 – inner membrane solute transporter

Mup1 - high-affinity methionine/cysteine permease

Mup3 – low-affinity methionine/cysteine permease

NCR - nitrogen catabolite repression

Ncs1 – neuronal calcium sensor 1

Nde2 – pyridine nucleotide-disulfide oxidoreductase, a homologue of *S. cerevisiae*

Ndh1- mitochondria external NADH dehydrogenase

 $NF-\kappa B$ – nuclear factor kappa-light-chain-enhancer of activated B cells

Nha1 – Na⁺/H⁺ antiporter induced for acidoresistance

Nic1 – nickel transporter

Nrg1/2 - putative C2H2-type transcription factor controlled by cAMP to regulate capsule production

NSR - nitrosative stress regulatory groups of genes

OMSR – osmotic stress response/regulatory groups of genes

Opt1 – oligopeptide transporter

OSR - oxidative stress response/regulatory groups of genes

Pbx – parallel β -helix repeat protein

Pck1 – phosphoenolpyruvate carboxykinase 1

Pcl12 - alternative cyclin

Pho4 – phosphate signalling pathway (PHO) activator

Pho81 - cyclin-dependent kinase (Cdk) inhibitor with conserved lysine surface cluster in the SPX domain

Pho84, Pho85, Pho89, Pho840 – high-affinity phosphate transporter/permease

Pka – cAMP-dependent protein kinase A catalytic subunit

Pkc1 – protein kinase *c*

Pqp1 – pro-Qsp1 protease 1 (a secreted serine protease)

Ptp – protein tyrosine kinase – Ptp1 and Ptp2 (a major negative repressor of Hog1 hyperphosphorylation)

Ptp1 – polyol transporter protein 1

Ptr – peptide transporter

Pub1 – polyadenylated RNA-binding protein

Puf4 – PUmilio-FBF (FEM-3 binding factor) homology domain family 4 of *m*RNA binding proteins very similar to yeast Mpt5. Together with **Lhp1** and **Pbp1** are calcineurin (Crz1) target genes for virulence and stress survival. They belong to P-bodies/stress granules

Pum1 – translational repressing RNA-binding protein

Rad17 - Radiation-sensitive cell cycle checkpoint protein associated with Mec1

Rad24 - Radiation-sensitive protein that promotes the binding of Rad17-Mec3-Ddc1 complex to DNA

Ras1 – GTPase involved in G-protein signalling in the adenylate cyclase-activating pathway

Rho – Rho-GTPase protein with C-terminal CAAX box motifs for prenylation and membrane localisation, a homologue of mammalian Rac proteins

Rim101 - cell wall transcriptional regulator

SCAP – SREBP cleavage-activating protein (same as Scp1p)

Sch9 – AGC/AKT protein kinase

Sec – exocyst complex component

Sec13 - COPII protein

Sec16 - COPII vesicle coat protein

Sec27 – coatomer β-subunit

Sec28 - COPI protein

Sec4 – secretory vesicle-associated Rab GTPase

Sec53 – phosphomannomutase:glycosylation secretory proteins

Sec61 – γ -subunit of protein translocation complex for misfolded proteins out of ER

Skn1 – β-glucan synthesis-associated protein putatively tagged glucosidase

Smt1 – sphingolipid C9 methyltransferase

Snf1 - sucrose non-fermenting serine/threonine-protein kinase 1 complex (homologue of AMPK in mammals)

Snf3 – glucose permease

Snf4 - Snf1 kinase complex with CBS binding domain

Snf7 – sucrose non-fermenting protein 7 – a key regulator of the endosomal sorting complex required for transport (ESCRT)-III for the formation and cargo of multivesicular bodies (MVB). It is a vacuolar-sorting protein

Soa1 – putative sulfonate/sulfur transporter

Sod1 – copper/zinc cytoplasmic superoxide dismutase

Sod2 – mitochondrial manganese superoxide dismutase

SRE – stress response elements

SREBP – mammalian sterol regulatory element-binding protein (Sre gene in fungus)

Ssk1 – sensory transduction histidine kinase

Ssn8/Srb11 – C-type cyclin protein associating with specific cyclin-dependent kinase (Cdk) like Cdk8

Ssn801 – cyclin C

Ssr – specific-stress regulatory genes

Suc2 – sucrose- and raffinose-degrading enzyme, invertase

Sul1/Sul2 – sulfate transporter/permease

Sur2 – sphinganine/sphingosine C-4 hydroxylase – an ER enzyme converting sphinganine to phytosphingosine; sterol desaturase

Swe102 – a homologue of *S. cerevisiae* Swe1 checkpoint protein kinase that regulates yeast morphogenesis by inhibiting Cdk via phosphorylation, an anti-stress mechanism in the budding yeast

Sxi1 α – sex inducer 1α for cell identity factor in MAT α similar to Sxi2**a** in MAT**a**

Tah18 - NAPDH-dependent diflavin reductase

Tao3 – RAM (Regulation of Ace2p activity and cellular Morphogenesis) is a cell polarity signalling network protein

Tco - two-component system

Tps1 – trehalose-6-phosphate synthase, catalysing the conversion of UDP-glucose and glucose-6-phosphate to trehalose-6-phosphate. UDP-galactose is an epimer of UDP-glucose, which is reported as a poor substrate and inhibitor of Tps1

Tps2 – trehalose-6-phosphate phosphatase

Trp1 – phosphoribosyl anthranilate isomerase in tryptophan biosynthesis

Trp2 – anthranilate synthase component I – ASCO-I of chorismate aminase

Trp3 – anthranilate synthase component II – ASCOII of glutamine amidotransferase/phosphoribosyl anthranilate isomerase/indole glycerol phosphate synthase

Trp4 – anthranilate phosphoribosyltransferase (APRT)

Trp5 – tryptophan synthase

Tsa1 – thiol-specific antioxidant thioredoxin peroxidase (thioredoxin-dependent peroxide reductase)

Usv101 - master regulator of C. neoformans pathogenesis and virulence, a C2H2 yeast transcription factor orthologue

Uve1 – endonuclease for base/nucleotide excision repair pathway in UV-DNA damage

Uxs1 – UDP-xylose synthase/UDP-glucuronate decarboxylase 1

Vad1 - virulence-associated DEAD-Box RNA helicase-encoding protein, a member of the RCK/p54 subfamily of RNA

Vcx1 - Ca²⁺/H⁺ antiporter

Vph1 – vacuolar/vesicular H⁺-ATPase proton pump

Vps1 – vacuolar protein sorting regulator, dynamin

Vps17 – a membrane coat complex retromer subunit Vps5/Snx1

Vps29/Vps36 – retrograde transporter (yeast vacuolar protein sorting-associated protein 29)

Vps34 – phosphatidylinositol-3-kinase (PI3K)

Vps4 – AAA+-type ATPase

Vrk1 – a homologue of vaccinia-related serine/threonine kinase induced by DNA damage

Yah1 – 2Fe-2S iron-sulfur cluster binding domain homologue of S. cerevisiae

Yor1- yeast oligomycin resistance

Ypc - alkaline phytoceramidase

Ypd – phosphotransfer protein/phosphorelay intermediate protein