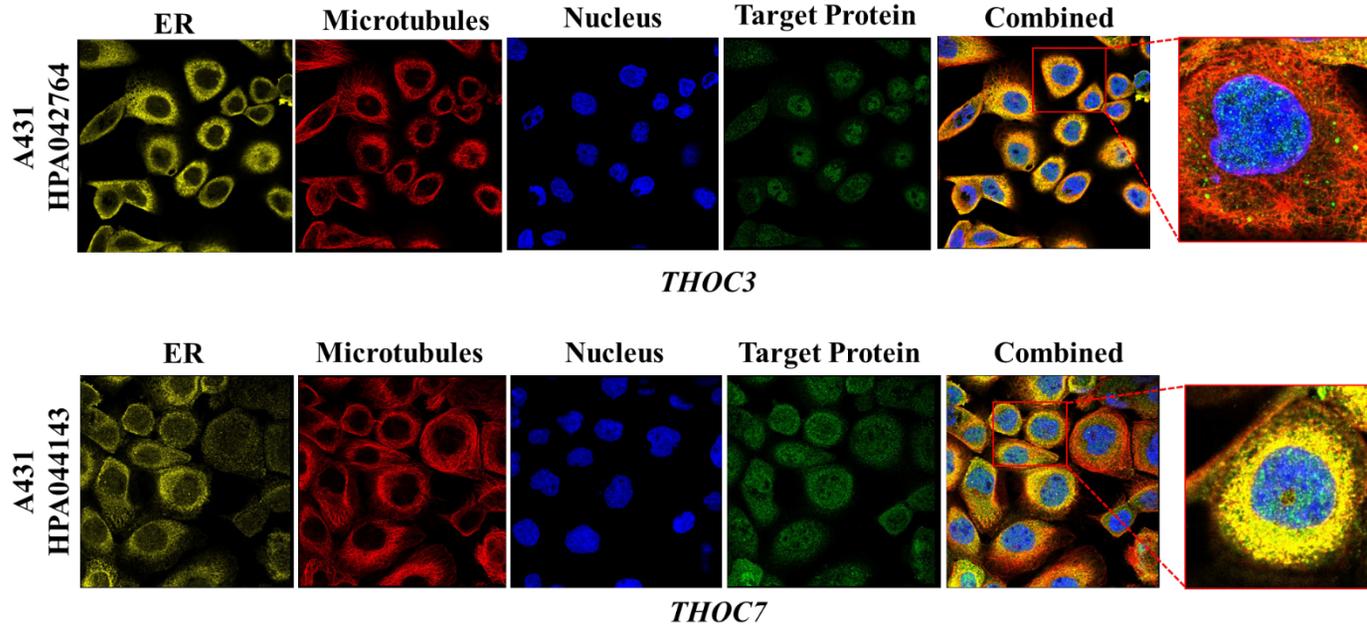
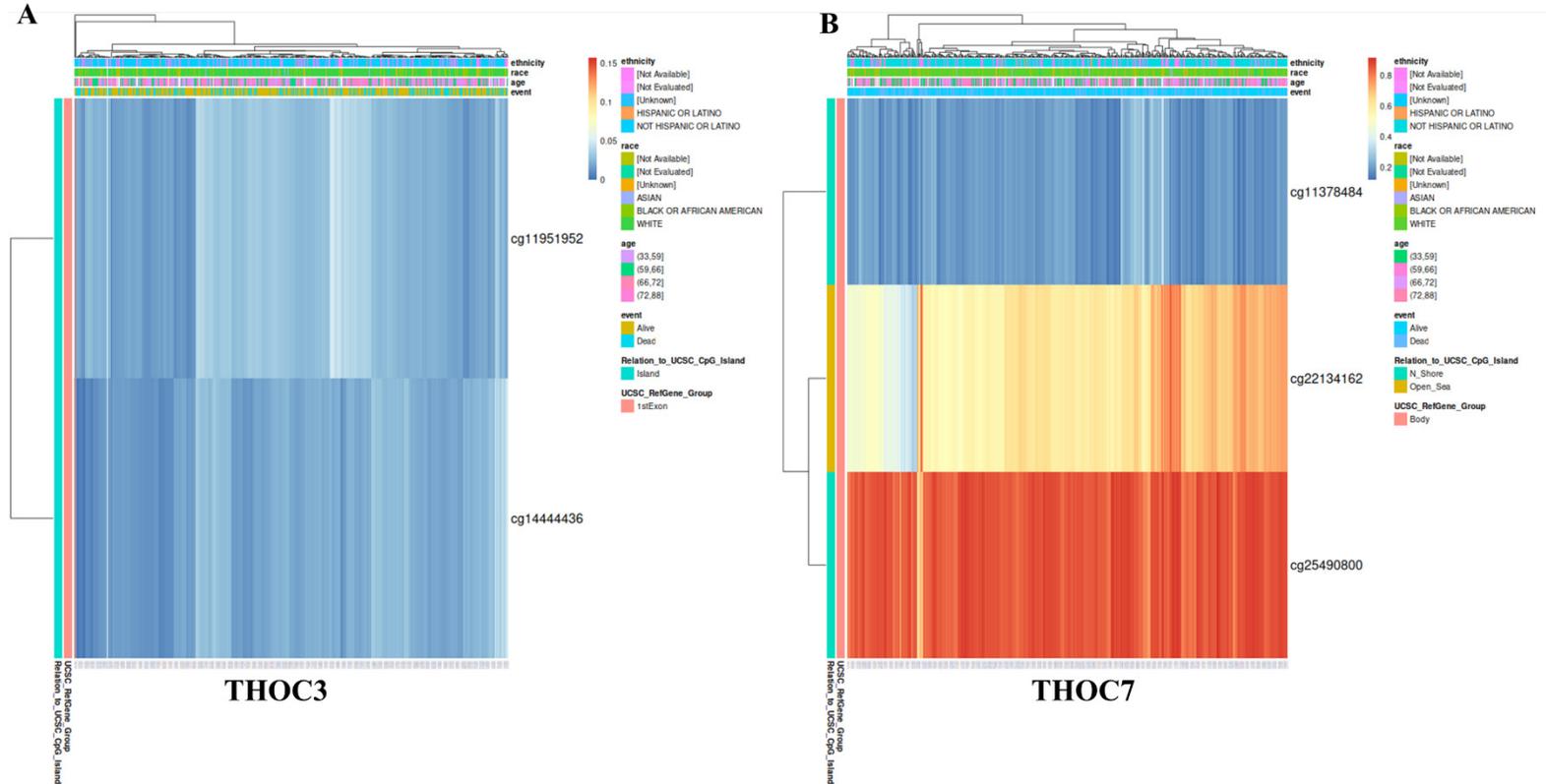


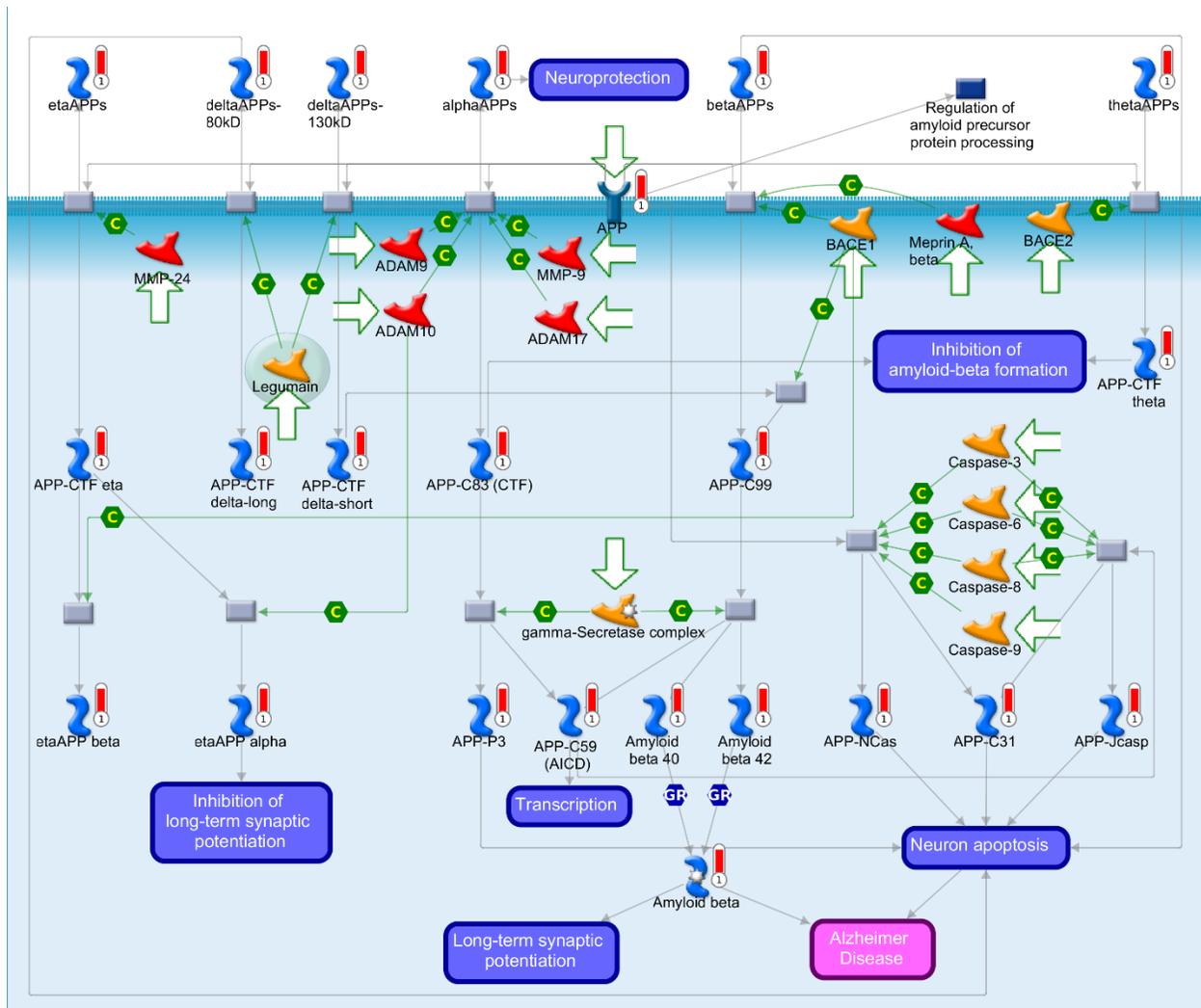
Supplementary Figure 1: Expression of THOC family genes across LUAD clinical stages and assessment of cohort-driven batch effects. (A) Percentage of variance explained by the top six principal components derived from principal component analysis (PCA) of combined TCGA-LUAD tumor and GTEx normal lung transcriptomic datasets, illustrating the relative contribution of each component to overall variance. (B) PCA plot (PC1 vs PC2) of combined TCGA-LUAD tumor samples and GTEx normal lung tissues prior to batch correction. Samples are colored by cohort, revealing clear cohort-driven separation, indicative of batch effects between TCGA and GTEx datasets. (C–H) Violin plots showing mRNA expression levels of THOC1 (A), THOC2 (B), THOC3 (C), THOC5 (D), THOC6 (E), and THOC7 (F) across pathological stages I–IV in lung adenocarcinoma (LUAD) patients from the TCGA cohort. Each violin represents the distribution of expression values within a given clinical stage. White dots indicate median expression levels, and black vertical bars denote the interquartile range. One-way ANOVA was used to evaluate differences in expression across stages; F statistics and corresponding p-values are shown in each panel. No statistically significant stage-dependent differences were observed for THOC family members, although THOC3 displayed a modest, non-significant trend toward higher expression in advanced stages.



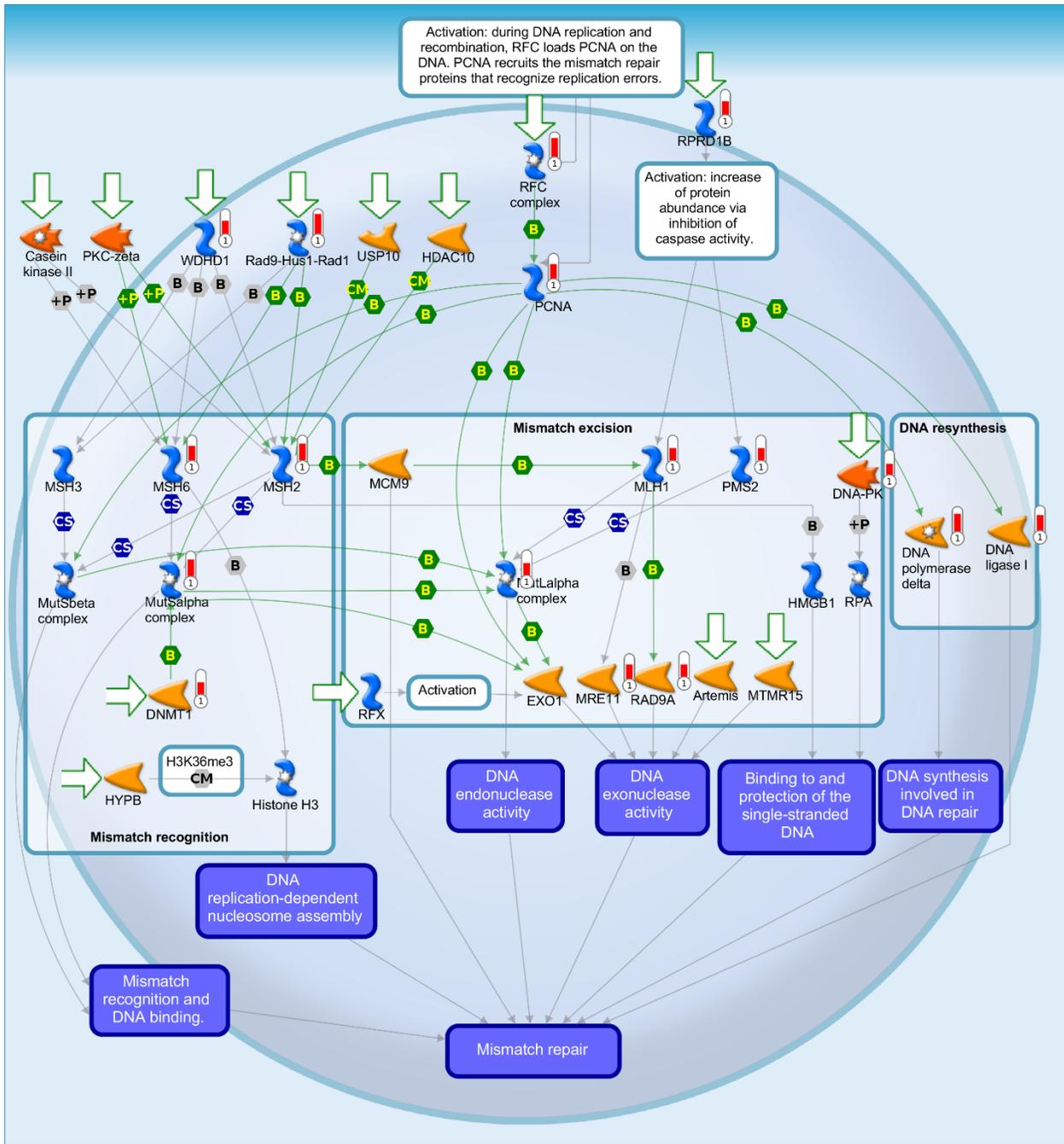
Supplementary Figure 2. Immunofluorescence validation of THOC3 and THOC7 subcellular localization in an EGFR-high epithelial model. Immunofluorescence staining was used to assess the subcellular localization of THOC3 and THOC7. THOC3 showed predominantly cytoplasmic and membranous distribution, whereas THOC7 was mainly localized to the nucleus, consistent with immunohistochemical findings in LUAD tissues. A431 cells were employed as an epithelial reference model due to their high EGFR expression, a key molecular feature of LUAD, and their well-defined cellular morphology, enabling clear visualization of EGFR-associated nucleocytoplasmic compartmentalization of THOC3 and THOC7.



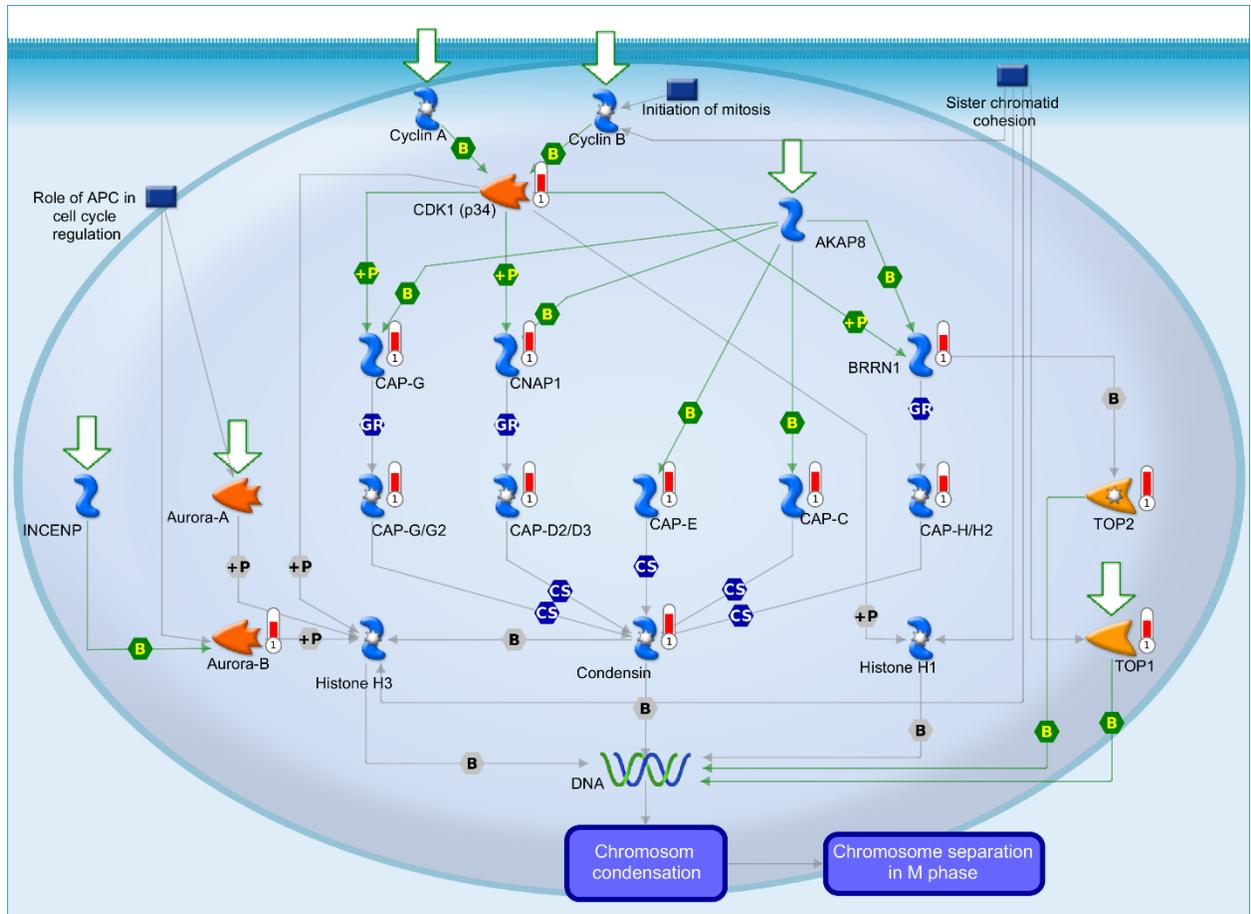
Supplementary Figure 3. Differential DNA methylation patterns of THOC3 and THOC7 in LUAD.(A) Heatmap showing methylation status of CpG sites associated with THOC3 (cg11951952 and cg1444436) across LUAD patient samples from TCGA. Predominantly hypomethylated regions are observed, suggesting epigenetic activation.(B) Heatmap showing CpG methylation levels for THOC7 (cg11378484, cg22134162, cg25490800). THOC7 exhibits broad hypomethylation across multiple CpG islands and gene body regions. Annotations for patient ethnicity, race, age, and survival status are displayed at the top of each heatmap.



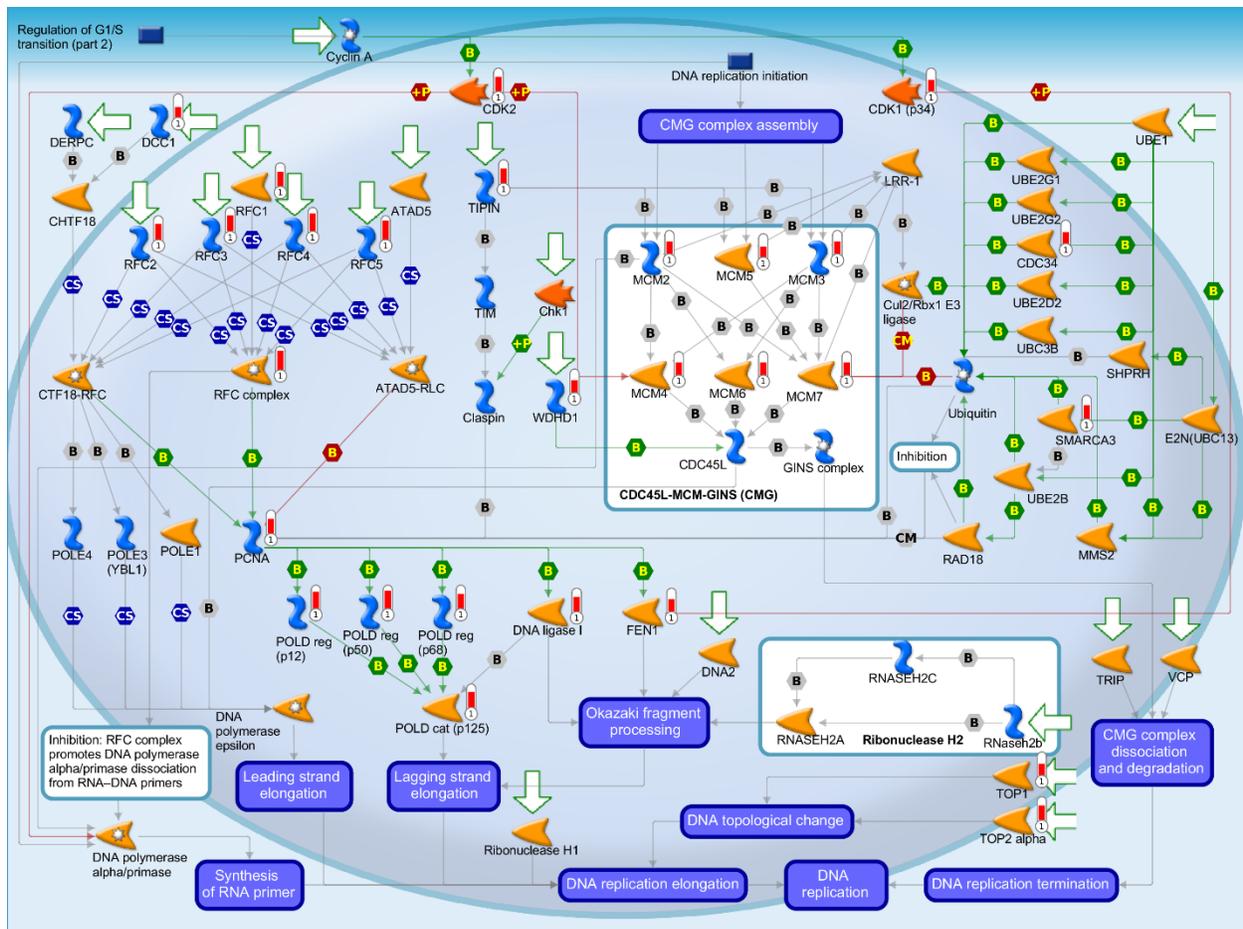
Supplementary Figure 4. Enrichment of the pathway Protein folding and maturation: amyloid precursor protein processing (schema) in LUAD for THOC3.



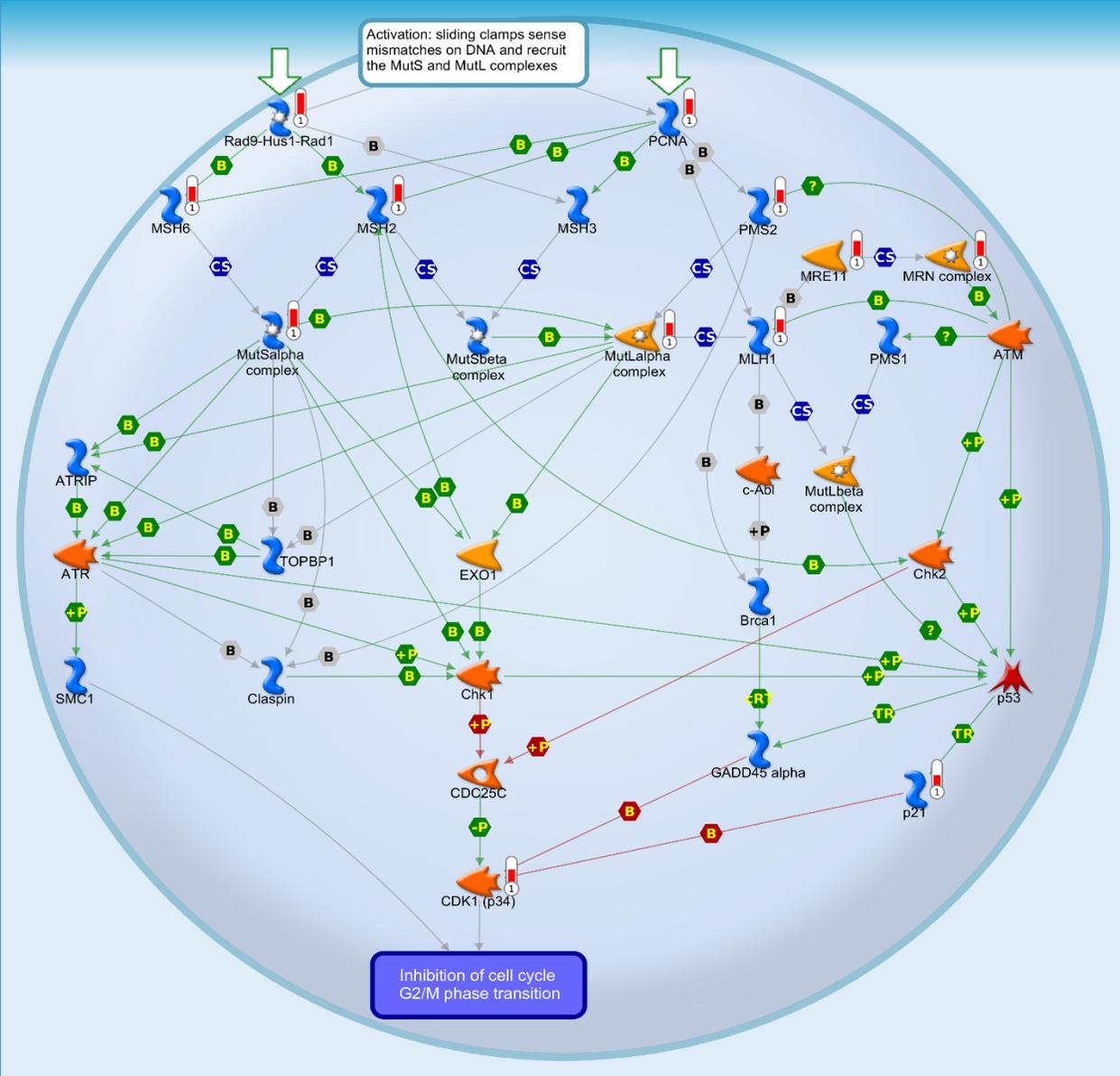
Supplementary Figure 5. Enrichment of the pathway DNA damage: mismatch repair in LUAD for THOC3.



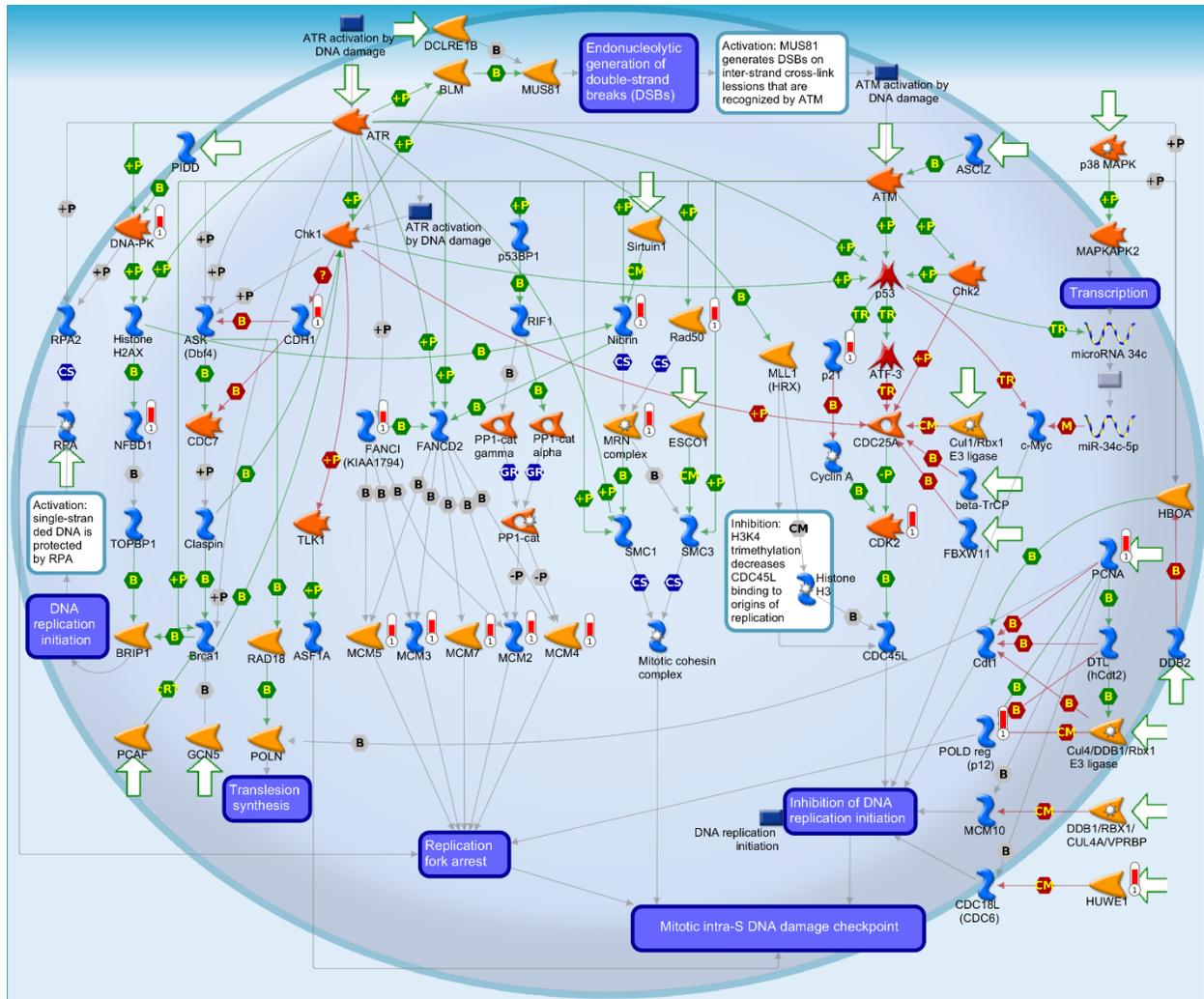
Supplementary Figure 6. Enrichment of the pathway Cell cycle: chromosome condensation in prometaphase in LUAD for THOC3.



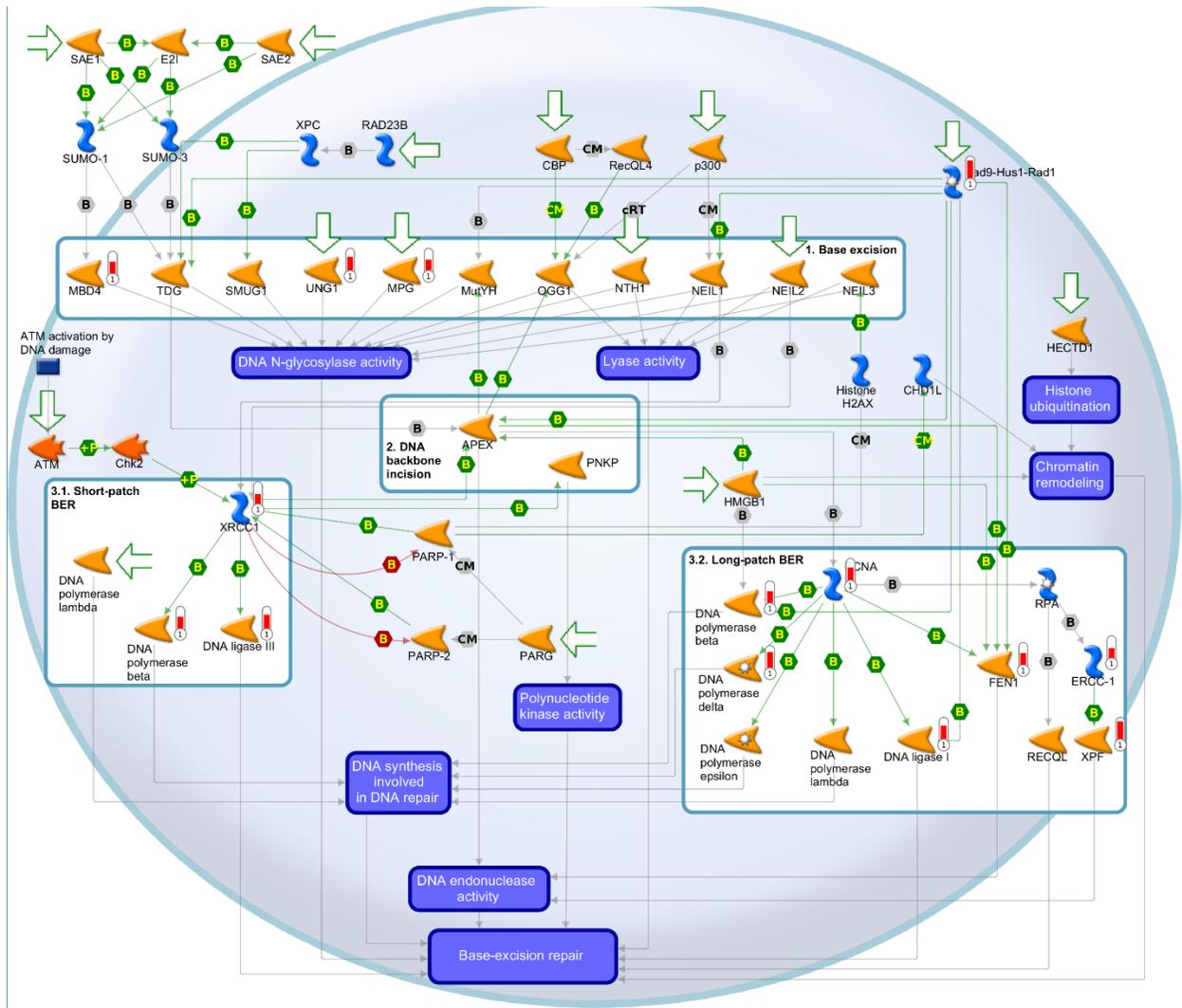
Supplementary Figure 7. Enrichment of the pathway Cell cycle: DNA replication initiation in LUAD for THOC3.



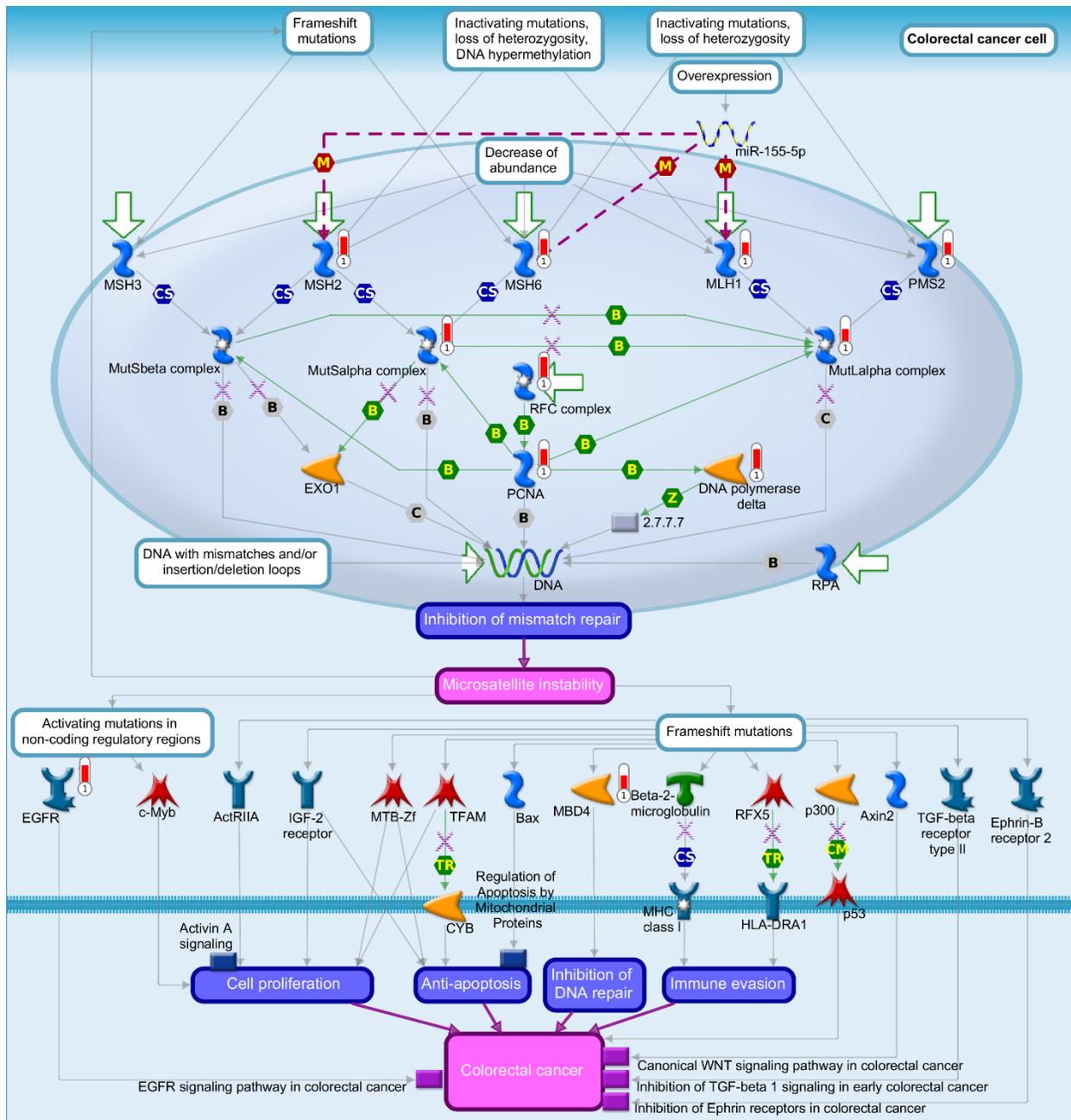
Supplementary Figure 8. Enrichment of the pathway DNA damage: G2 checkpoint in response to DNA mismatches in LUAD for THOC3.



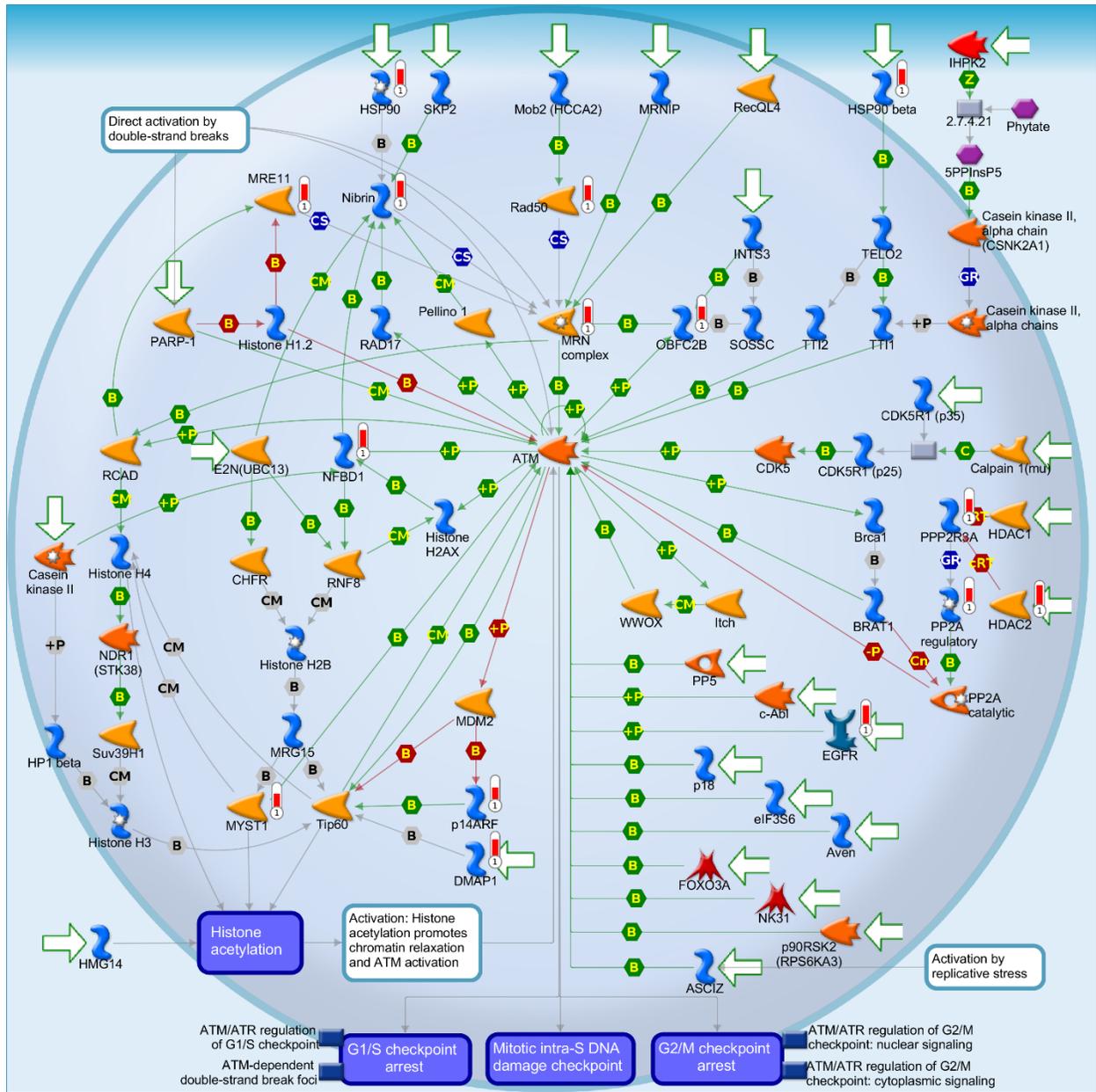
Supplementary Figure 9. Enrichment of the pathway DNA damage: intra-S-phase checkpoint in LUAD for THOC3.



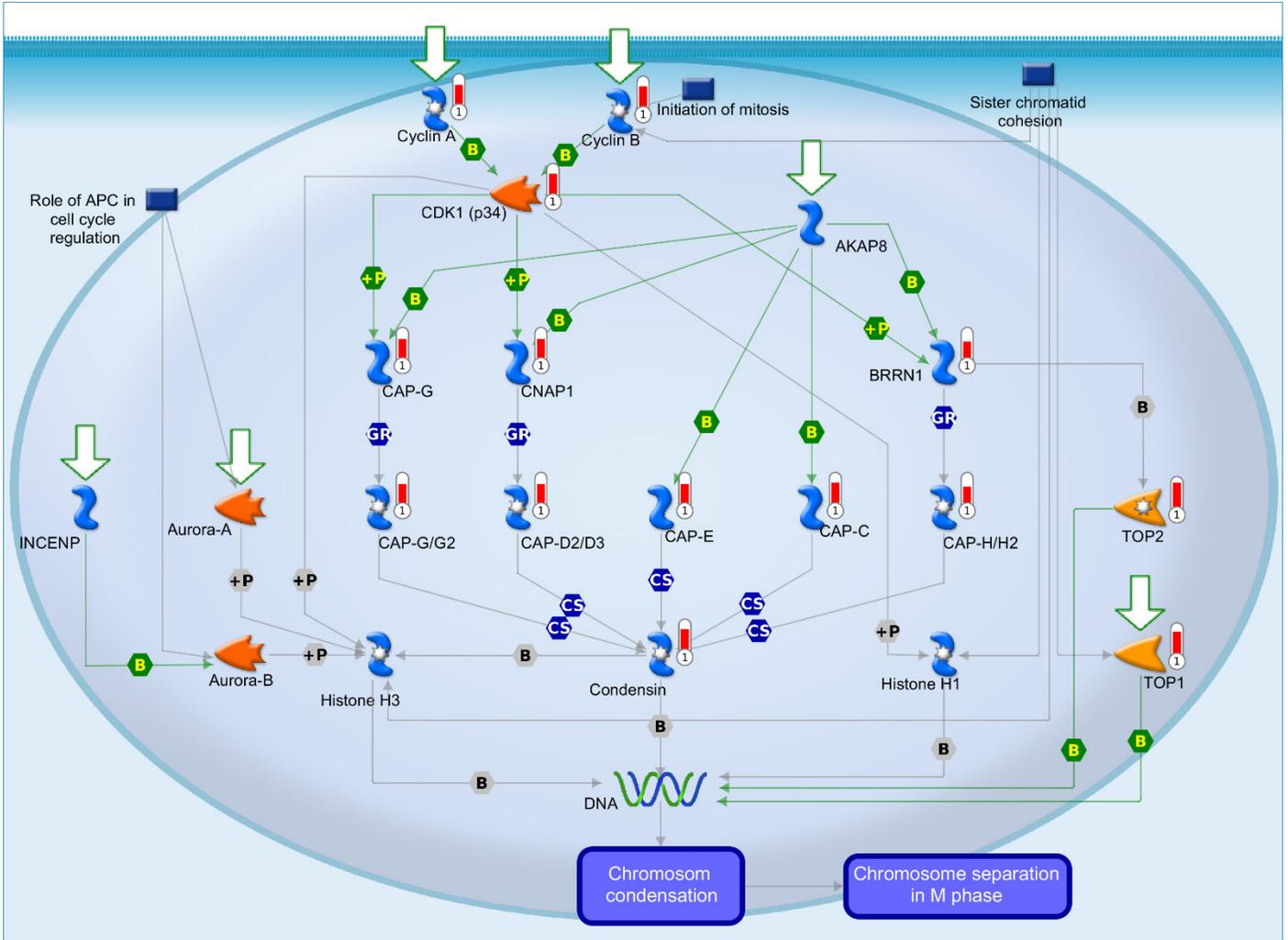
Supplementary Figure 10. Enrichment of the pathway DNA damage: base excision repair (BER) in LUAD for THOC3.



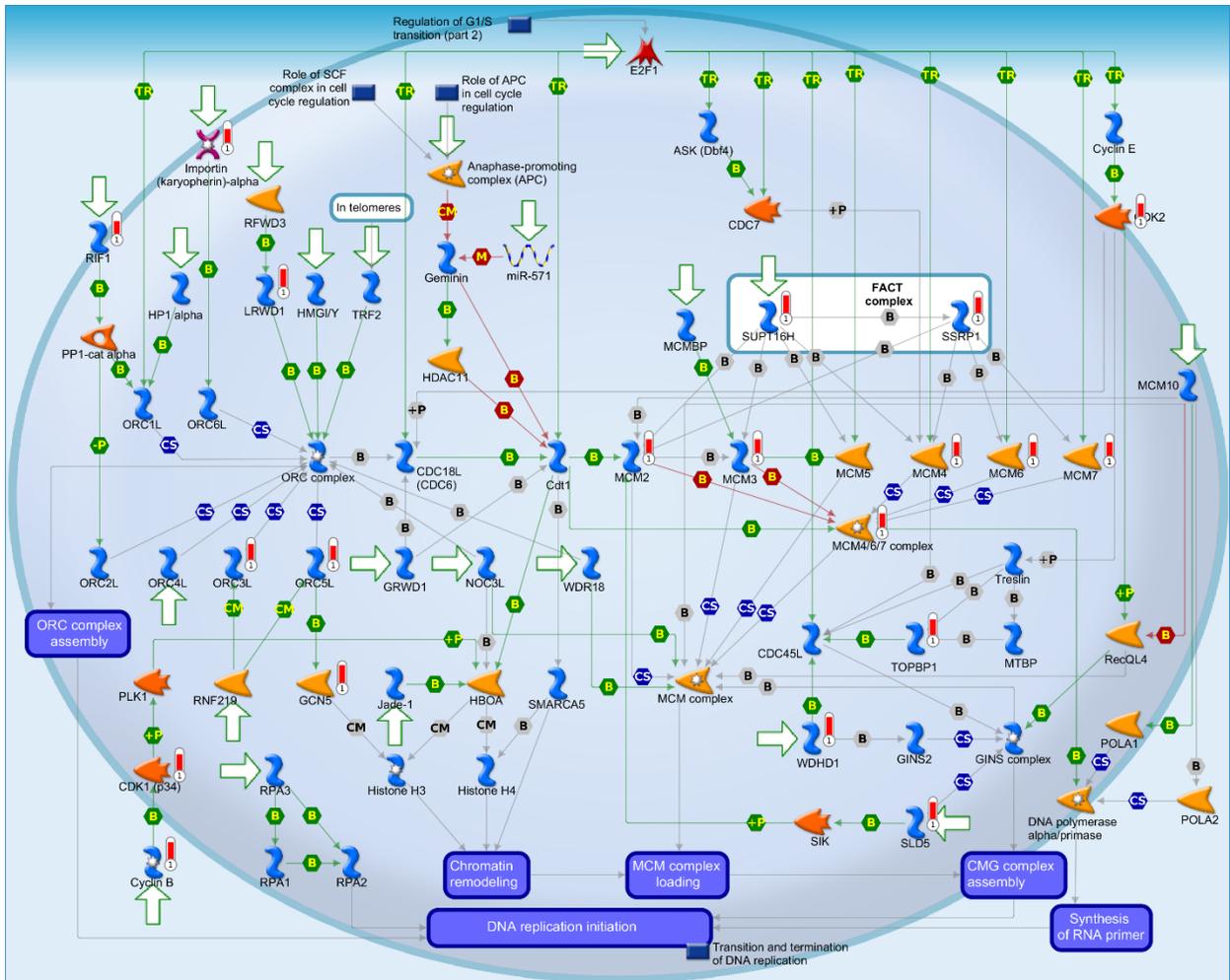
Supplementary Figure 11. Enrichment of the pathway Microsatellite instability (MSI) in colorectal cancer in LUAD for THOC3.



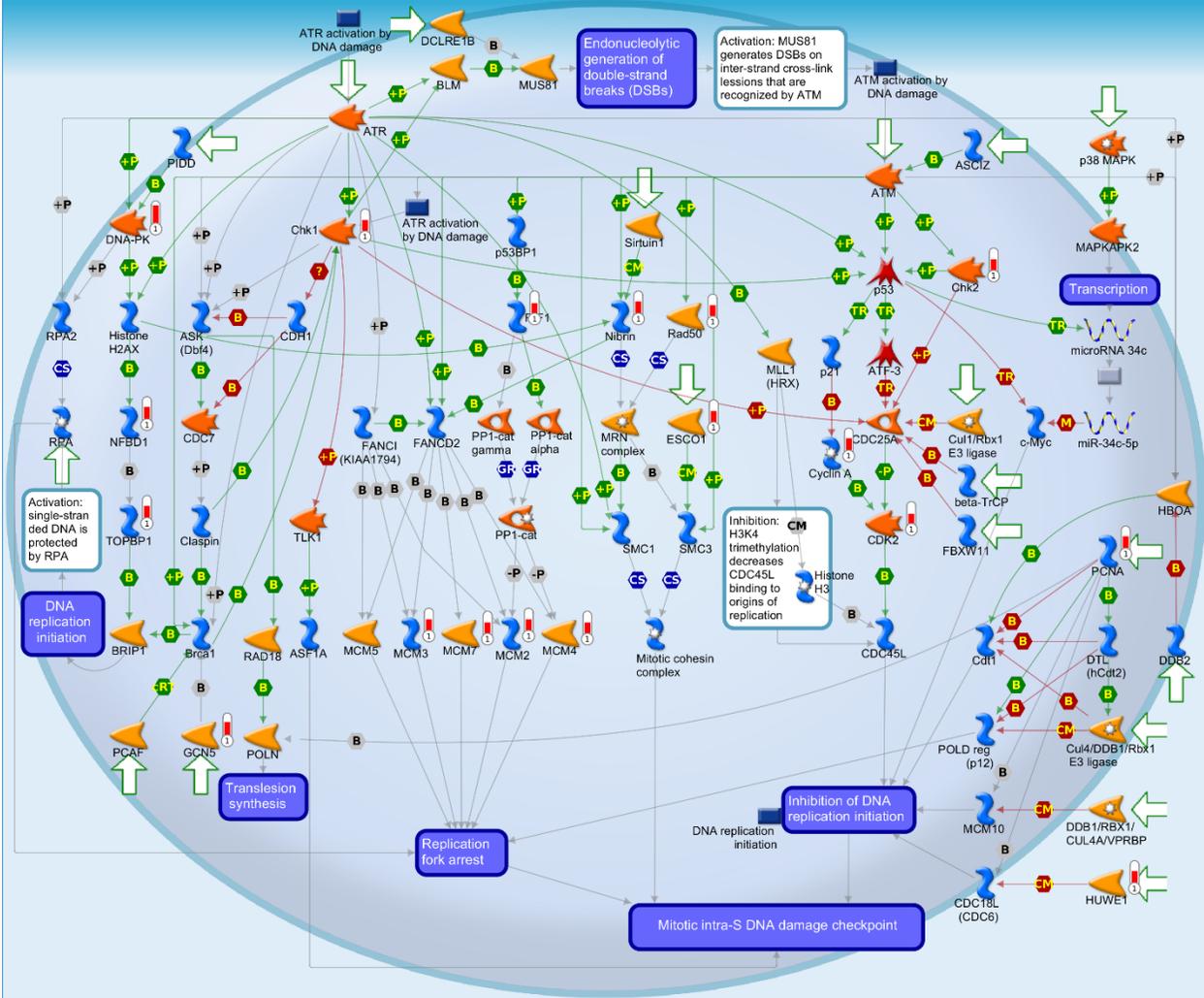
Supplementary Figure 12. Enrichment of the pathway DNA damage: ATM activation by DNA damage in LUAD for THOC3



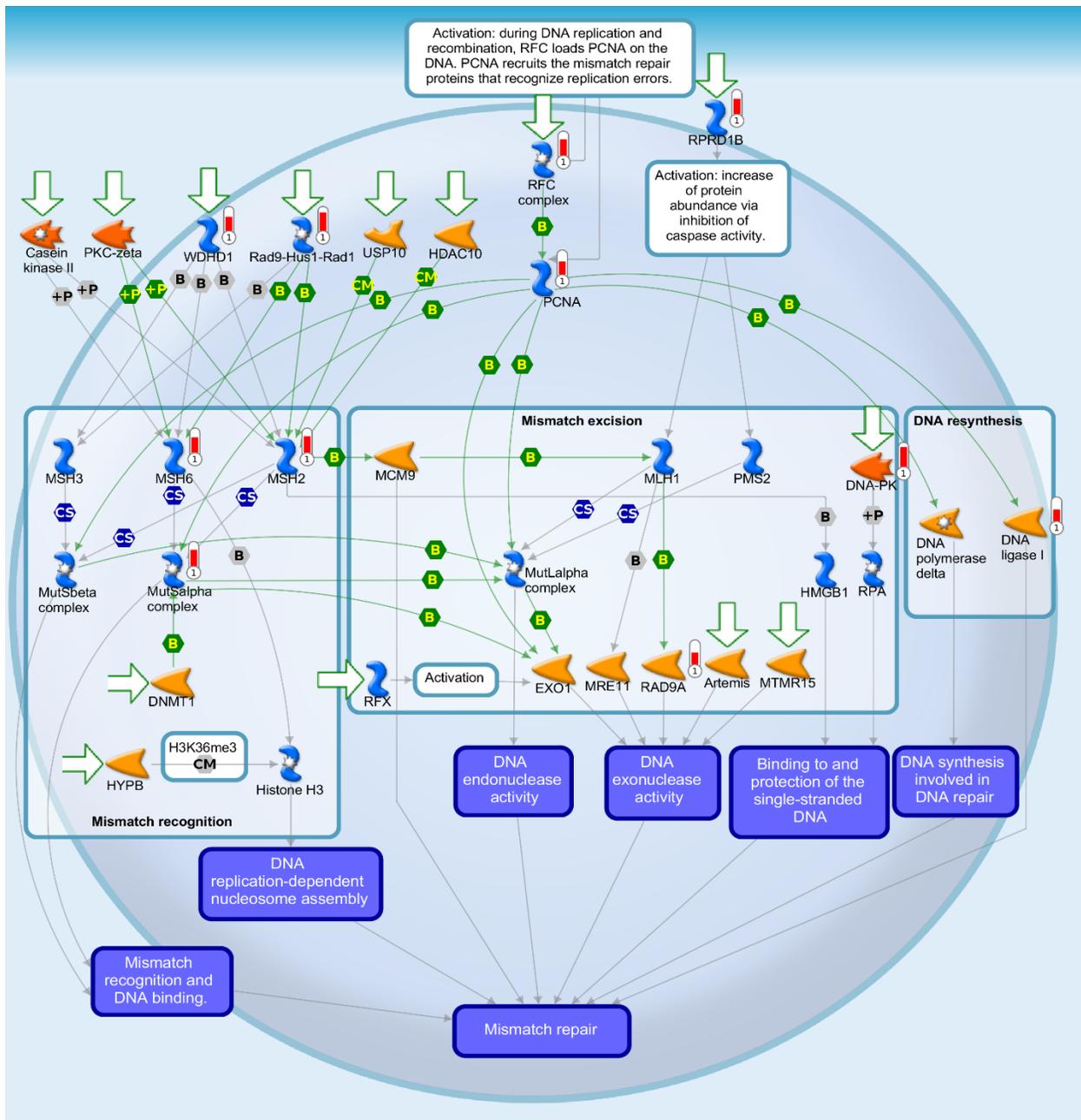
Supplementary Figure 13. Enrichment of the pathway Cell cycle: chromosome condensation in prometaphase in LUAD for THOC7.



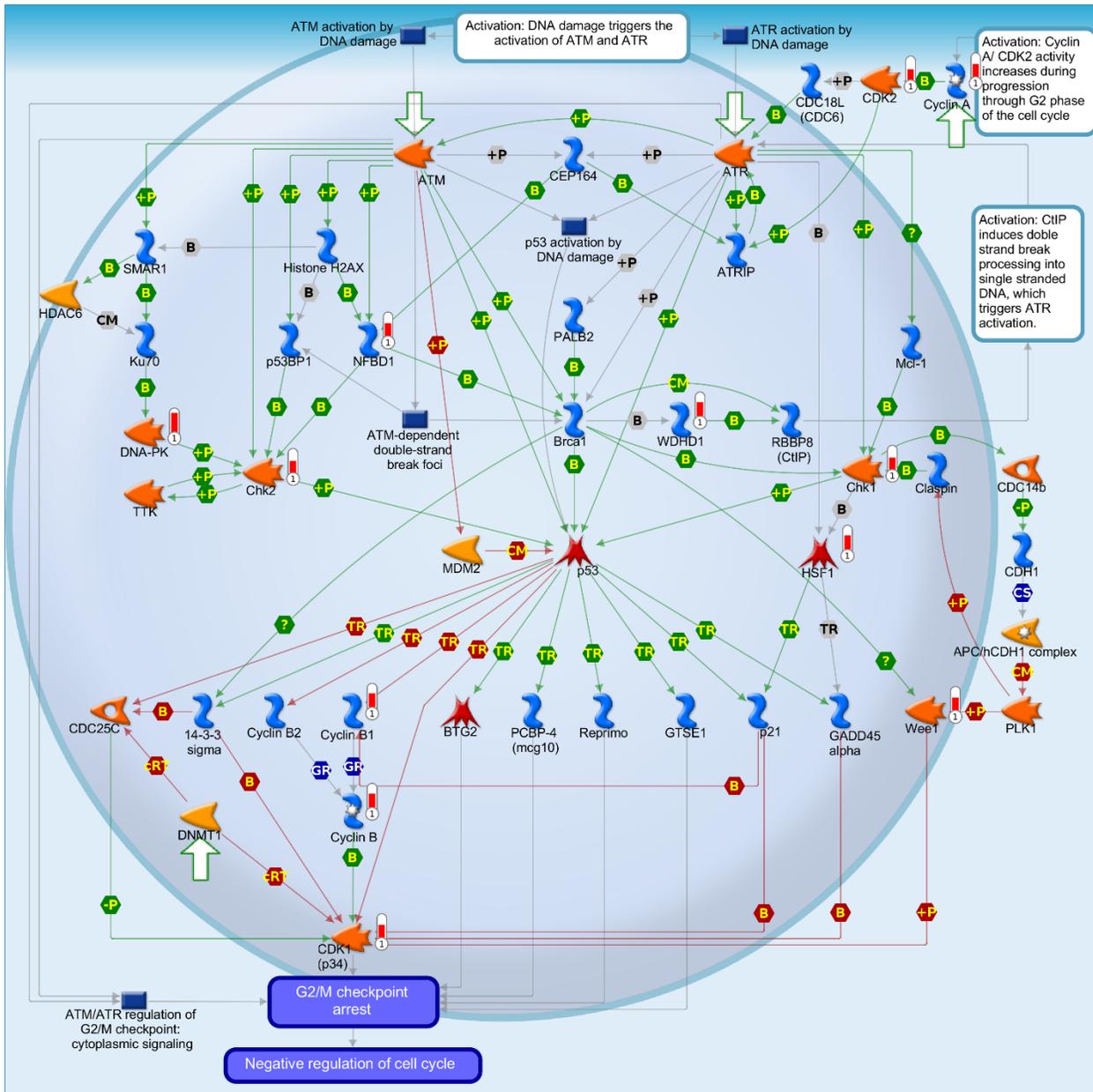
Supplementary Figure 14. Enrichment of the pathway Cell cycle: DNA replication initiation in LUAD for THOC7.



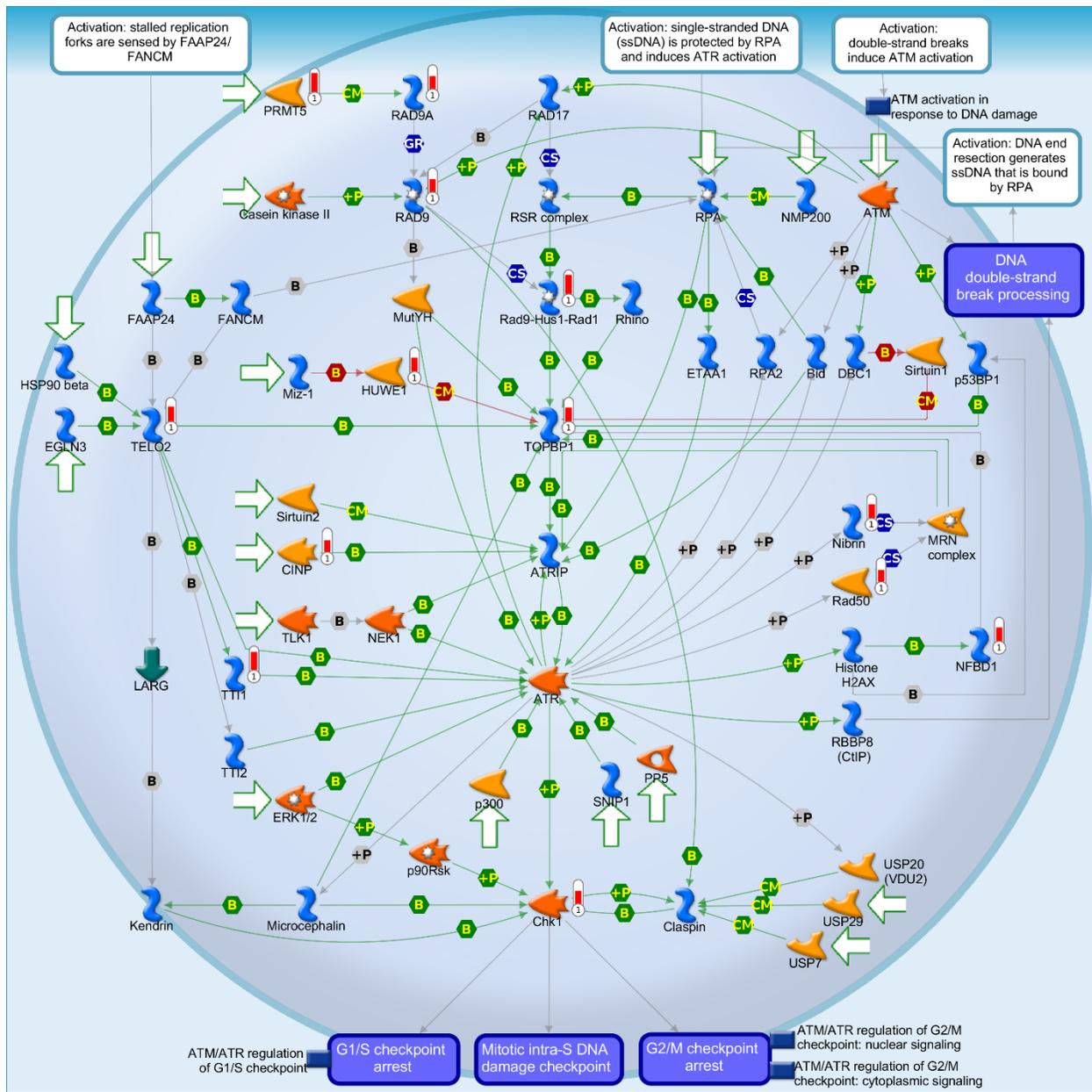
Supplementary Figure 15. Enrichment of the pathway DNA damage: intra-S-phase checkpoint in LUAD for THOC7.



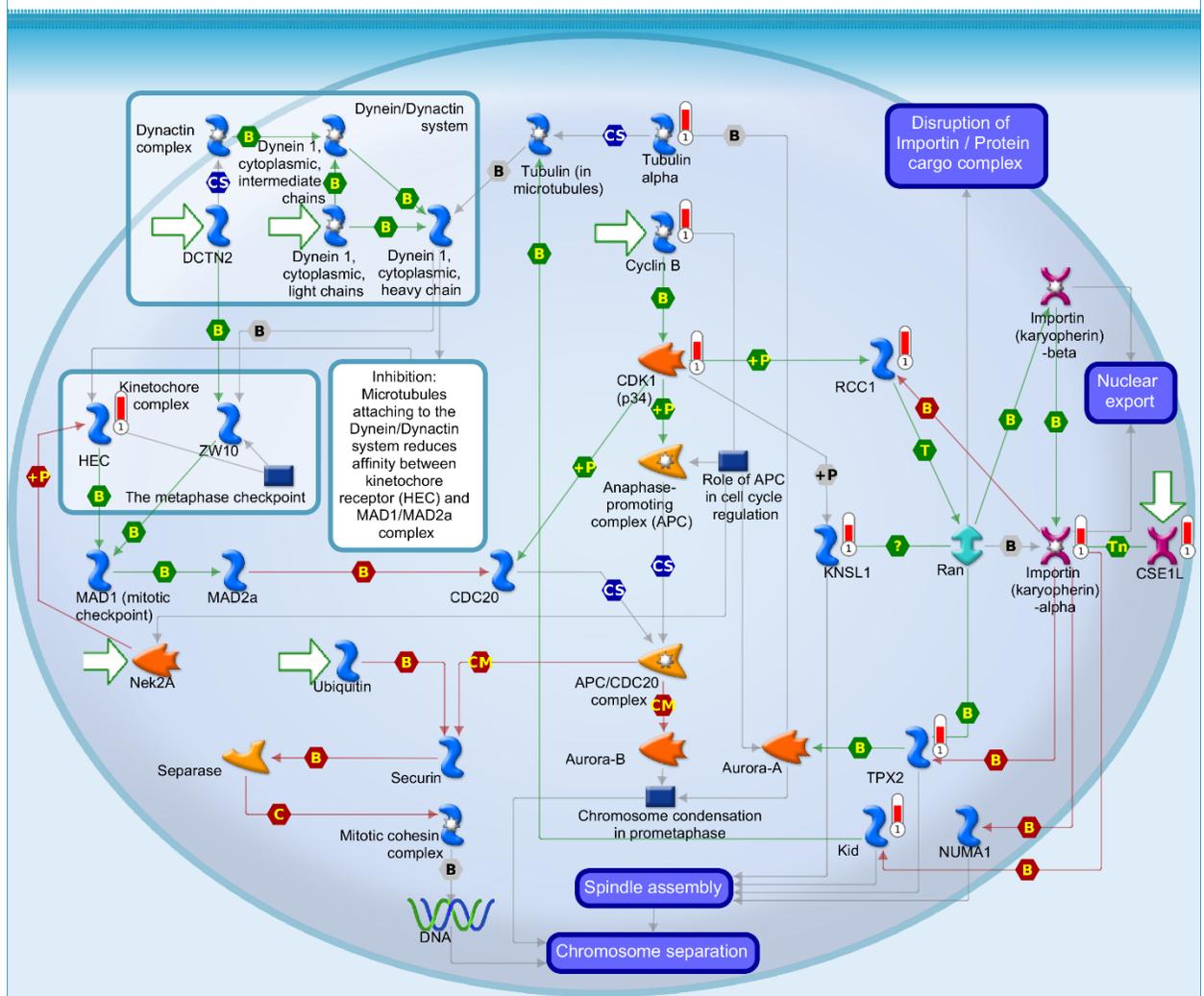
Supplementary Figure 16. Enrichment of the pathway DNA damage: mismatch repair in LUAD for THOC7.



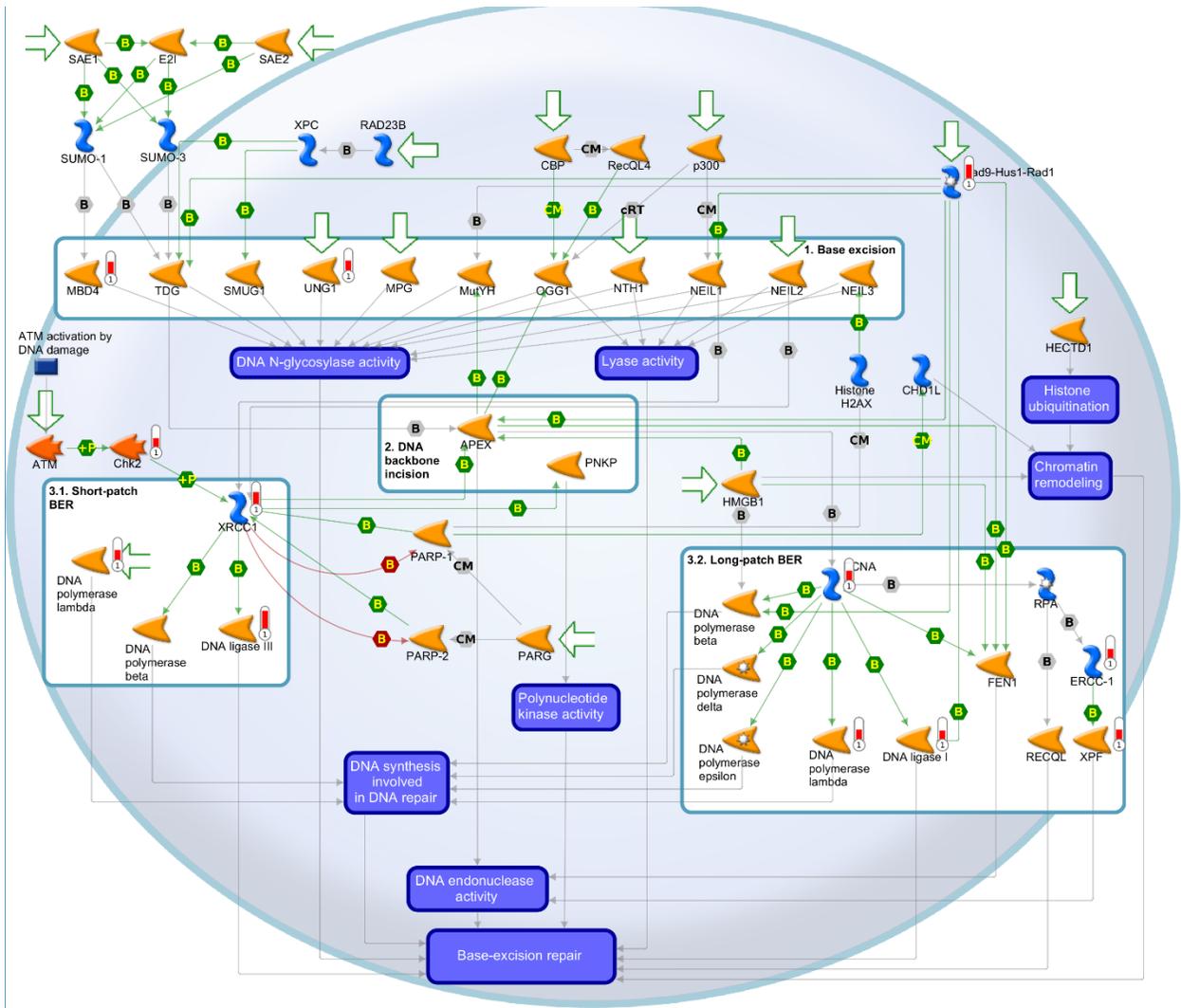
Supplementary Figure 17. Enrichment of the pathway DNA damage: ATM/ATR regulation of G2/M checkpoint nuclear signaling in LUAD for THOC7.



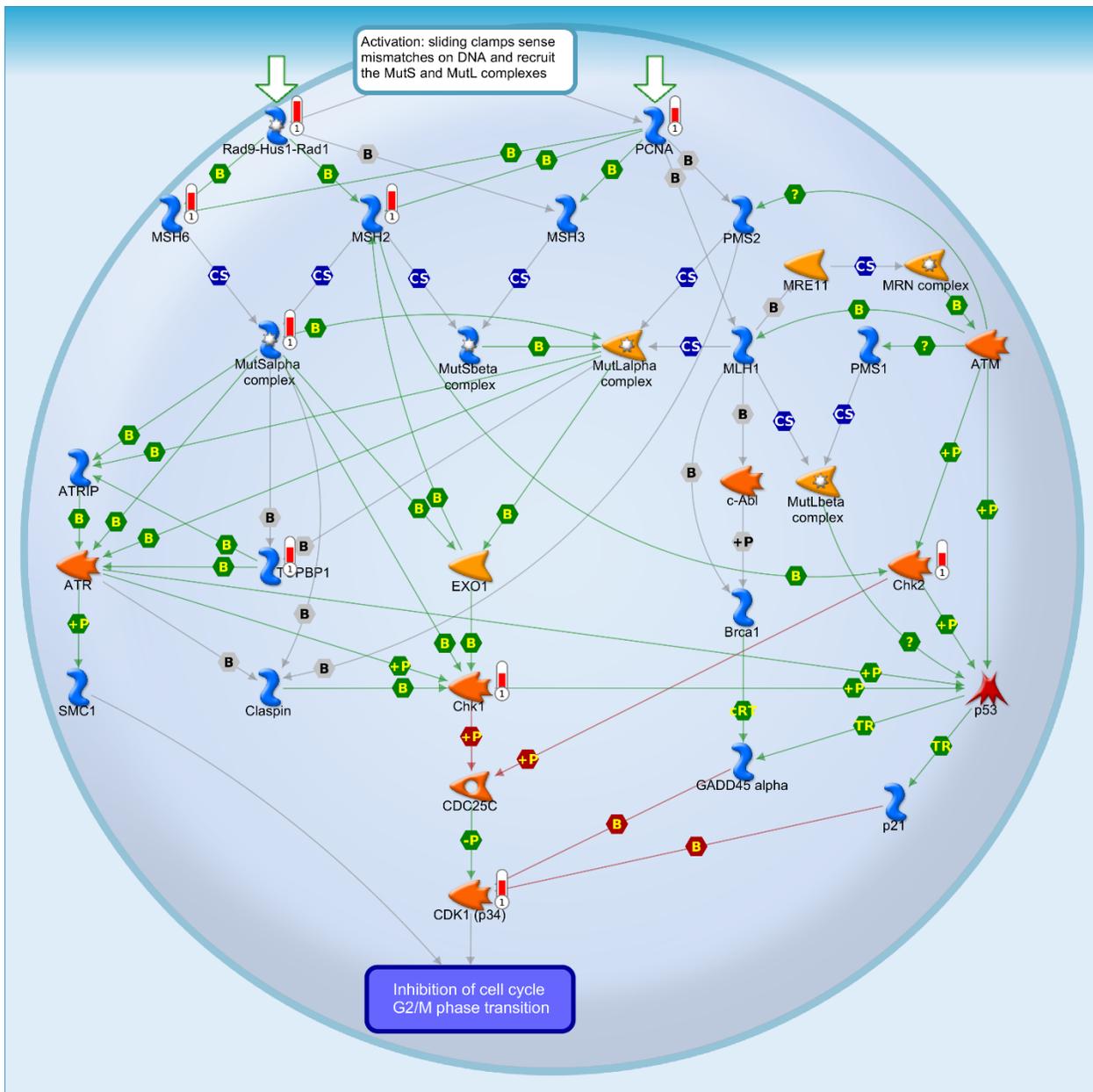
Supplementary Figure 18. Enrichment of the pathway DNA damage: ATR activation by DNA damage in LUAD for THOC7.



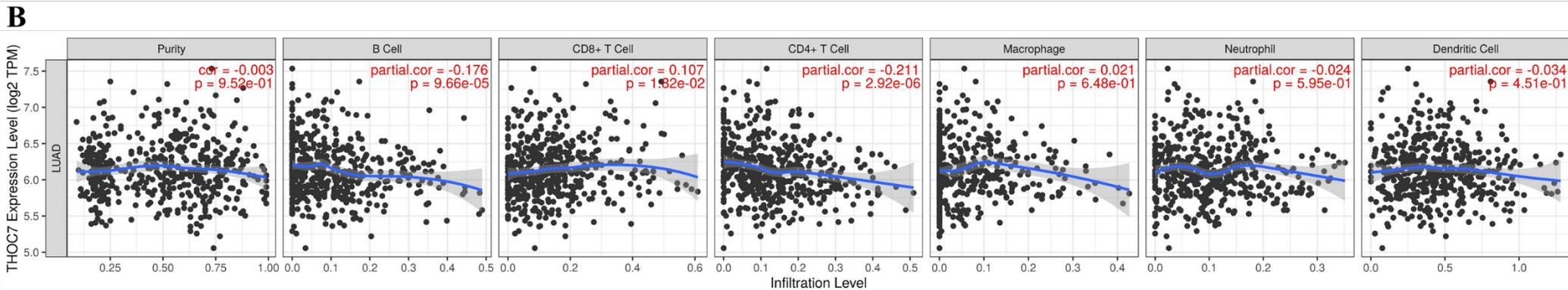
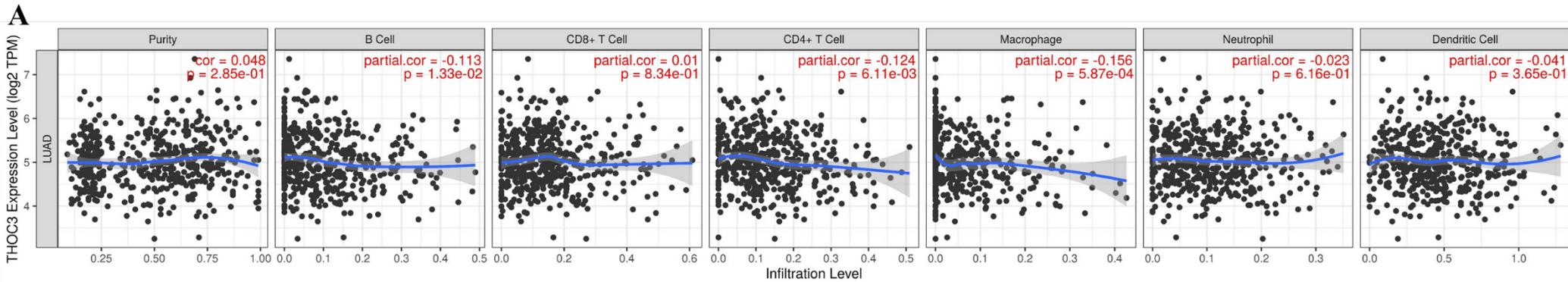
Supplementary Figure 19. Enrichment of the pathway Cell cycle: spindle assembly and chromosome separation in LUAD for THOC7.



Supplementary Figure 20. Enrichment of the pathway DNA damage: base excision repair (BER) in LUAD for THOC7.



Supplementary Figure 21. Enrichment of the pathway DNA damage_G2 checkpoint in response to DNA mismatches in LUAD for THOC7.



Supplementary Figure 22. Immune infiltration profiles of THOC3 and THOC7 in LUA (A–B) Scatterplots depicting correlations between THOC3 expression (C) and THOC7 expression profiles with immune cell infiltration in LUAD. Analysis was performed across multiple immune subtypes, including B cells, CD8⁺ T cells, CD4⁺ T cells, macrophages, neutrophils, and dendritic cells.

Table 1: Pathway analysis of co-expressed genes with THOC3 from the MetaCore database

S No	Maps	pValue	Network Objects from Active Data
1	Cell cycle_DNA replication: elongation and termination	1.213E-24	RFC4, TOP2 alpha, WDHD1, PCNA, MCM6, MCM3, RFC1, RFC2, MCM7, TOP1, RFC3, MCM2, POLD reg (p50), FEN1, POLD cat (p125), DNA ligase I, MCM5, RFC5, RFC complex, MCM4, TIPIN, POLD reg (p68), SMARCA3, DCC1, POLD reg (p12), CDK1 (p34), CDC34, CDK2
2	Protein folding and maturation_Amyloid precursor protein processing (schema)	4.485E-21	APP-C99, APP-C31, APP-C83 (CTF), APP-CTF delta-short, etaAPP alpha, APP-NCas, Amyloid beta 40, APP-CTF delta-long, betaAPPs, APP, deltaAPPs-80kD, APP-CTF eta, thetaAPPs, APP-P3, alphaAPPs, Amyloid beta, etaAPPs, etaAPP beta, APP-C59 (AICD), deltaAPPs-130kD, APP-CTF theta, APP-Jcasp, Amyloid beta 42
3	DNA damage_Mismatch repair	2.864E-17	WDHD1, PCNA, MutSalpha complex, PMS2, DNMT1, MRE11, DNA polymerase delta, RAD9A, MSH2, DNA ligase I, RPRD1B, Rad9-Hus1-Rad1, MLH1, RFC complex, MSH6, MutLalpha complex, DNA-PK
4	Cell cycle_Chromosome condensation in prometaphase	1.091E-14	CAP-H/H2, Condensin, CAP-C, CNAP1, CAP-D2/D3, Aurora-B, TOP1, TOP2, BRRN1, CAP-G, CAP-G/G2, CAP-E, CDK1 (p34)
5	Cell cycle_DNA replication initiation	3.741E-14	WDHD1, Importin (karyopherin)-alpha, MCM6, MCM3, SSRP1, SMARCA5, MCM7, RPA1, MCM4/6/7 complex, POLA1, MCM2, MCM complex, MCM5, RPA3, SLD5, MCM4, SUPT16H, LRWD1, CDK1 (p34), CDK2
6	DNA damage_G2 checkpoint in response to DNA mismatches	1.027E-10	PCNA, MutSalpha complex, PMS2, MRE11, MSH2, p21, Rad9-Hus1-Rad1, MRN complex, MLH1, MSH6, MutLalpha complex, CDK1 (p34)
7	DNA damage_Intra S-phase checkpoint	4.860E-10	PCNA, CDH1, MCM3, Rad50, NFB1, MCM7, MCM2, MCM5, p21, MRN complex, MCM4, FANCI (KIAA1794), Nibrin, HUWE1, POLD reg (p12), DNA-PK, CDK2
8	DNA damage_Base excision repair	2.506E-09	PCNA, UNG1, MPG, XRCC1, DNA polymerase delta, DNA polymerase beta, XPF, ERCC-1, FEN1, DNA ligase I, Rad9-Hus1-Rad1, MBD4, DNA ligase III
9	Microsatellite instability in colorectal cancer	8.796E-09	PCNA, MutSalpha complex, PMS2, DNA polymerase delta, MSH2, MBD4, MLH1,

			RFC complex, MSH6, EGFR, MutLalpha complex
10	DNA damage_ATM activation by DNA damage	1.190E-08	PP2A regulatory, Rad50, MRE11, p14ARF, NFB1, DMAP1, HDAC2, HSP90, PPP2R3A, OBFC2B, MRN complex, Nibrin, EGFR, HSP90 beta, MYST1
11	Microsatellite instability in gastric cancer	1.886E-08	PCNA, MutSalpha complex, PMS2, MSH2, MLH1, RFC complex, p16INK4, MSH6, MutLalpha complex
12	DNA damage_Double-strand break repair via homologous recombination	1.284E-07	FIGNL1, WDHD1, PPP4C, MRE11, SMARCA5, NFB1, RPA1, MRN complex, WDR79, BRD9, Nibrin, CCDC98 (Abraxas), RAP80, CDK1 (p34), CDK2
13	DNA damage_ATR activation by DNA damage	1.307E-06	Rad50, PRMT5, NFB1, RAD9A, Rad9-Hus1-Rad1, MRN complex, Nibrin, HUWE1, HSP90 beta, USP7, RAD9
14	DNA damage_ATM-dependent double-strand break foci	1.921E-06	SMARCA5, NFB1, ENL, HBXAP, MRN complex, BAT3, Mi-2 beta, Nibrin, USP7, CCDC98 (Abraxas), RAP80, KDM2A
15	DNA damage_Nucleotide excision repair	1.921E-06	PCNA, RFC1, XRCC1, DNA polymerase delta, XPF, ERCC-1, DNA ligase I, MPP11, RFC complex, DNA ligase III, SUPT16H, USP7
16	Cell cycle_The metaphase checkpoint	2.335E-06	PMF1, BUB3, DSN1, Aurora-B, ZW10, BUB1, CDCA1, Zwilch, BUBR1
17	Transcription_Negative regulation of HIF1A function	3.136E-06	SART1, MCM3, p14ARF, MCM7, MCM2, MCM5, Elongin B, HSP90, HSP70, HSPA4, HSP90 beta, LAMP2
18	Abnormalities in cell cycle in small cell lung cancer (SCLC)	3.847E-06	PCNA, p14ARF, Aurora-B, CKS1, p21, p16INK4, CDK1 (p34), CDK2
19	Cell cycle_Sister chromatid cohesion	3.861E-06	PCNA, TOP1, RFC3, PDS5, RFC complex, Stromalins 1/2, DCC1
20	dCTP/dUTP metabolism	6.694E-06	AK2, POLA1, DNA polymerase beta, POLD reg (p50), POLD cat (p125), RRM2, TK1, POLG reg, POLD reg (p68), Small RR subunit, AK1, POLD reg (p12)
21	Transcription_Sin3 and NuRD in transcription regulation	7.471E-06	Mi-2, MBD3, HDAC2, RBBP4 (RbAp48), Mi-2 beta, SMRT, Sin3B, RAR-alpha/RXR-beta, RARalpha
22	Cell cycle_Role of APC in cell cycle regulation	8.617E-06	BUB3, CDH1, Aurora-B, CKS1, BUB1, CDK1 (p34), CDK2, BUBR1
23	Cell cycle_Spindle assembly and chromosome separation	1.104E-05	Importin (karyopherin)-alpha, RCC1, KNSL1, Aurora-B, Tubulin alpha, ZW10, CSE1L, CDK1 (p34)
24	dATP/dITP metabolism	1.677E-05	8ODP, AK2, POLA1, DNA polymerase beta, POLD reg (p50), POLD cat (p125), Adenosine kinase, RRM2, POLG reg, POLD reg (p68), Small RR subunit, AK1, POLD reg (p12)
25	Aberrant B-Raf signaling in melanoma progression	8.890E-05	Rictor, Aurora-B, B-Raf, RHEB2, p21, Nicastrin, HES1, SRp55, CDK1 (p34)

26	DNA damage_ATM/ATR regulation of G2/M checkpoint: nuclear signaling	1.015E-04	WDHD1, CDH1, DNMT1, NFBD1, p21, DNA-PK, CDK1 (p34), CDK2
27	CFTR folding and maturation (normal and cystic fibrosis)	1.195E-04	Calnexin, HSP90 alpha, HSP70, Aha1, HSP90 beta, FKBP8
28	ATP/ITP metabolism	2.799E-04	AK2, POLR1A, Adenosine kinase, RPB5, ENTPD6, RRM2, HPRT, RPB7.0, Small RR subunit, 5'-NT1A, ADAR1, RPB8, AK1
29	Immune response_ETV3 affect on CSF1-promoted macrophage differentiation	3.485E-04	DDX20, HDAC2, SMRT, PRIM2A, CDK1 (p34)
30	Brca1 and Brca2 in breast cancer	3.485E-04	Rad50, MSH2, MRN complex, MLH1, Nibrin
31	DNA damage_Classical NHEJ mechanism of DSBs repair	3.538E-04	PAXIP1L, NFBD1, Aprataxin, TARDBP (TDP43), HDAC2, MRN complex, DNA-PK
32	TTP metabolism	4.664E-04	POLA1, UDP, DNA polymerase beta, POLD reg (p50), POLD cat (p125), TK1, POLG reg, POLD reg (p68), POLD reg (p12)
33	Possible regulation of HSF-1/chaperone pathway in Huntington's disease	5.765E-04	HSP90 alpha, HSP90, PLA2, HSP70, HSP90 beta
34	Tau dysregulation in Alzheimer disease	5.984E-04	Caspase-2, CAPON, LCMT1, PP2A regulatory, OGT (GlcNAc transferase), TARDBP (TDP43), APP, HSP90, APP-C59 (AICD), Amyloid beta 42
35	HSP70 and HSP40-dependent folding in Huntington's disease	1.345E-03	HSP90 alpha, HSP90, HSP70, SGTA, HSP90 beta
36	dGTP metabolism	1.401E-03	8ODP, POLA1, DNA polymerase beta, POLD reg (p50), POLD cat (p125), POLG reg, POLD reg (p68), POLD reg (p12)
37	Stem cells_Cytotoxic effect of temozolomide on glioblastoma cells	1.933E-03	MutSalph complex, MSH2, p21, MSH6, MutLalpha complex
38	DNA damage_p53 activation by DNA damage	2.079E-03	PP2A regulatory, AATF (Che-1), p21, TTC5 (Strap), USP7, MYST1, PP2C gamma, DNA-PK
39	Transcription_Role of heterochromatin protein 1 (HP1) family in transcriptional silencing	2.152E-03	Mi-2, DNMT1, SETDB1, MBD3, HDAC2, CDK1 (p34)
40	Cell cycle_Senescence activation pathways	2.233E-03	p14ARF, RHEB2, HSP90, p21, p16INK4, WISp39, CDK2
41	The role of aberrations in CDKN2 locus and CDK4 in familial melanoma	2.289E-03	p14ARF, p21, p16INK4, EGFR, MIG6
42	Mitogenic action of Estradiol / ESR1 (nuclear) in breast cancer	2.289E-03	CHD8, p21, NCOA3 (pCIP/SRC3), LRP16, CDK2

43	Maturation and migration of dendritic cells in skin sensitization	2.450E-03	MHC class II alpha chain, HLA-DRB1, MHC class II, HLA-DRB5, MHC class II beta chain, HLA-DRB
44	Cell cycle_Role of Cul1/Rbx1 E3 ligase in cell cycle regulation	2.689E-03	CKS1, p21, CDK1 (p34), CDC34, CDK2
45	Transcription_Epigenetic regulation of gene expression	3.056E-03	DNMT1, PRMT5, SETDB1, SMCX, JMJD1A, HDAC2, MYST1
46	DNA damage_ATM/ATR regulation of G1/S checkpoint	3.137E-03	PCNA, PP2A regulatory, NFBD1, p21, MRN complex, CDK2
47	Apoptosis and survival_Regulation of apoptosis by mitochondrial proteins	3.240E-03	Caspase-2, RAD9A, NOR1, PP2C, Aif, Fis1, TIMM8A, RAD9, Cofilin, CDK2
48	Development_H3K36 demethylation in stem cell maintenance	4.147E-03	p14ARF, M33, p21, p16INK4
49	Regulation of metabolism_ChREBP signaling	4.918E-03	PYC, PP2A regulatory, OGT (GlcNAc transferase), ACLY, SCD, SMRT, G6PT
50	Epigenetic alterations in ovarian cancer	6.644E-03	HNF1-beta, DNMT1, Aurora-B, HDAC2, p21, MLH1, p16INK4, SUZ12

Table 2: Pathway analysis of co-expressed genes with THOC7 from the MetaCore database

S No	Maps	pValue	Network Objects from Active Data
1	Cell cycle_DNA replication: elonaton and termination	2.684E-17	RFC4, TOP2 alpha, WDHD1, PCNA, MCM6, MCM3, RFC1, RFC2, Cyclin A, Chk1, MCM7, TOP1, RFC3, MCM2, DNA ligase I, RFC5, RFC complex, MCM4, TIPIN, SMARCA3, CDK1 (p34), CDC34, CDK2
2	Cell cycle_Chromosome condensation in prometaphase	7.157E-16	CAP-H/H2, Condensin, CAP-C, Cyclin A, CNAP1, CAP-D2/D3, TOP1, Cyclin B, TOP2, BRRN1, CAP-G, CAP-G/G2, CAP-E, CDK1 (p34)
3	Cell cycle_DNA replication initiation	1.916E-13	WDHD1, Importin (karyopherin)-alpha, MCM6, MCM3, SSRP1, RIF1, MCM7, ORC5L, Cyclin B, MCM4/6/7 complex, MCM2, TOPBP1, SLD5, MCM4, SUPT16H, ORC3L, LRWD1, CDK1 (p34), CDK2, GCN5
4	DNA damage_Intra S-phase checkpoint	2.224E-10	ESCO1, PCNA, Chk2, MCM3, Cyclin A, RIF1, Rad50, Chk1, NFB1, MCM7, MCM2, TOPBP1, MCM4, Nibrin, HUWE1, DNA-PK, CDK2, GCN5
5	DNA damage_Base excision repair	7.149E-09	PCNA, UNG1, Chk2, XRCC1, RAD23B, DNA polymerase beta, XPF, ERCC-1, DNA ligase I, Rad9-Hus1-Rad1, MBD4, DNA ligase III, DNA polymerase lambda
6	DNA damage_Mismatch repair	1.044E-08	WDHD1, PCNA, MutSalpha complex, RAD9A, MSH2, DNA ligase I, RPRD1B, Rad9-Hus1-Rad1, RFC complex, MSH6, DNA-PK
7	DNA damage_ATM/ATR regulation of G2/M checkpoint: nuclear signaling	4.252E-08	WDHD1, HSF1, Chk2, Cyclin A, Chk1, NFB1, Wee1, Cyclin B, Cyclin B1, DNA-PK, CDK1 (p34), CDK2
8	DNA damage_ATR activation by DNA damage	4.659E-08	TELO2, Rad50, Chk1, PRMT5, NFB1, RAD9A, C1NP, Rad9-Hus1-Rad1, TOPBP1, TTI1, Nibrin, HUWE1, RAD9
9	Cell cycle_Spindle assembly and chromosome separation	1.935E-07	Importin (karyopherin)-alpha, RCC1, KNSL1, HEC, Kid, Tubulin alpha, Cyclin B, TPX2, CSE1L, CDK1 (p34)
10	DNA damage_G2 checkpoint in response to DNA mismatches	8.830E-07	PCNA, MutSalpha complex, Chk2, Chk1, MSH2, Rad9-Hus1-Rad1, TOPBP1, MSH6, CDK1 (p34)
11	Transcription_Role of heterochromatin protein 1 (HP1) family in transcriptional silencing	1.403E-06	TIF1-beta, Mi-2, SETDB1, MBD3, HP1 beta, HDAC2, Cyclin A2, HP1 gamma, HP1, CDK1 (p34)

12	Transcription_Sin3 and NuRD in transcription regulation	1.794E-06	Mi-2, MBD3, p66beta, HDAC2, PSF, RBBP4 (RbAp48), Mi-2 beta, Sin3B, RAR-alpha/RXR-beta, RARalpha
13	DNA damage_Double-strand break repair via homologous recombination	2.276E-06	FIGNL1, WDHD1, PPP4C, RIF1, RMI1, NFB1, AUNIP, RecQ5, TOPBP1, WDR79, RAD54L, Nibrin, CDK1 (p34), CDK2
14	Cell cycle_Sister chromatid cohesion	6.856E-06	ESCO1, PCNA, TOP1, RFC3, PDS5, RFC complex, Stromalins 1/2
15	Abnormalities in cell cycle in small cell lung cancer (SCLC)	7.312E-06	PCNA, Cyclin A, p14ARF, CKS1, Cyclin B1, p16INK4, CDK1 (p34), CDK2
16	Microsatellite instability in colorectal cancer	2.626E-05	PCNA, MutSalpha complex, MSH2, MBD4, RFC complex, MSH6, EGFR, c-Myb
17	DNA damage_Nucleotide excision repair	2.774E-05	PCNA, RFC1, XRCC1, RAD23B, XPF, ERCC-1, DNA ligase I, MPP11, RFC complex, DNA ligase III, SUPT16H
18	Cell cycle_Initiation of mitosis	3.300E-05	Nucleolin, KNSL1, Wee1, Cyclin B1, MAT1, CDK1 (p34), CDK7
19	DNA damage_ATM activation by DNA damage	5.595E-05	TELO2, Rad50, p14ARF, NFB1, HP1 beta, HDAC2, HSP90, OBFC2B, TTI1, Nibrin, EGFR
20	Proteolysis_SUMOylation pathway	7.559E-05	TIF1-beta, MMS21, ZNF451, TRIM33, p14ARF, UHRF2, HDAC2, PIAS3
21	Possible regulation of HSF-1/ chaperone pathway in Huntington's disease	8.530E-05	HSF1, HSP90 alpha, HSP90, PLA2, HSP70, p23 co-chaperone
22	Cell cycle_Nucleocytoplasmic transport of CDK/Cyclins	1.044E-04	Importin (karyopherin)-alpha, Cyclin A, Cyclin B1, CDK1 (p34), CDK2
23	Development_Negative regulation of WNT/Beta-catenin signaling in the nucleus	1.050E-04	P15RS, NF-AT3(NFATC4), TRIM33, BCL9/B9L, Calcineurin A (catalytic), CHD8, TLE, HDAC2, RUVBL2, Kaiso, Frizzled, 14-3-3
24	Microsatellite instability in gastric cancer	1.136E-04	PCNA, MutSalpha complex, MSH2, RFC complex, p16INK4, MSH6
25	Development_H3K9 demethylases in pluripotency maintenance of stem cells	1.136E-04	RING2, TIF1-beta, SETDB1, JMJD1A, HP1, LBP9
26	Cell cycle_Role of 14-3-3 proteins in cell cycle regulation	1.136E-04	Chk2, Chk1, Wee1, HUS1, 14-3-3 zeta/delta, CDK1 (p34)
27	Cell cycle_Role of APC in cell cycle regulation	1.392E-04	Cyclin A, Kid, Cyclin B, CKS1, BUB1, CDK1 (p34), CDK2
28	DNA damage_ATM-dependent double-strand break foci	1.470E-04	RING2, TIF1-beta, NFB1, BAZ1A, HBXAP, BAT3, Mi-2 beta, Nibrin, HP1 gamma, GCN5
29	DNA damage_DNA-damage-induced responses	2.033E-04	Chk2, Chk1, NFB1, DNA-PK
30	Transcription_Negative regulation of HIF1A function	2.145E-04	SART1, MCM3, p14ARF, MCM7, MCM2, Elongin B, HSP90, HSP70, RUVBL2, Elongin C

31	Development_WNT/Beta-catenin signaling in the nucleus	4.360E-04	MED1, PYGO2, BCL9/B9L, TLE, Kaiso, Frizzled, UBR5, GCN5, RUVBL1
32	Development_Positive regulation of WNT/Beta-catenin signaling in the cytoplasm	4.778E-04	TGT, COX-2 (PTGS2), Bcl-9, BIG1, TGIF, 14-3-3 zeta/delta, Frizzled, CDK1 (p34), FAK1, 14-3-3
33	Cell cycle_Role of Cul1/Rbx1 E3 ligase in cell cycle regulation	5.802E-04	Chk1, Wee1, CKS1, CDK1 (p34), CDC34, CDK2
34	DNA damage_Classical NHEJ mechanism of DSBs repair	5.966E-04	RIF1, PAXIP1L, NFB1, Aprataxin, HDAC2, DNA polymerase lambda, DNA-PK
35	Development_Transcription factors in segregation of hepatocytic lineage	7.028E-04	HNF1-beta, p14ARF, CPSM, Activin A, p16INK4, Activin
36	Signal transduction_Glucocorticoid receptor signaling	7.852E-04	Keratin 17, MED1, TFB2M, HSP90, Karyopherin alpha 2, FKBP4, TFB1M, Oct-1
37	Development_Positive regulation of WNT/Beta-catenin signaling in the nucleus	9.686E-04	SOX4, BCL9/B9L, FOXK1, TLE, HDAC2, RUVBL2, NCOA2 (GRIP1/TIF2), Frizzled, UBR5
38	Cell cycle_Role of Nek in cell cycle regulation	1.007E-03	RCC1, HEC, Tubulin alpha, Cyclin B1, TPX2, CDK1 (p34)
39	CFTR folding and maturation (normal and cystic fibrosis)	1.631E-03	HSP90 alpha, Csp, HSP70, HSPBP1, p23 co-chaperone
40	Transcription_Mechanism of activation of the transcription of Retinoid-target genes	1.631E-03	MED1, CRABP2, NCOA3 (pCIP/SRC3), NCOA2 (GRIP1/TIF2), RARalpha
41	Cell cycle_The metaphase checkpoint	1.910E-03	SPBC25, CENP-B, HEC, BUB1, HP1 gamma, CDCA1
42	HSP70 and HSP40-dependent folding in Huntington's disease	1.977E-03	HSP90 alpha, HSP90, HSP70, SGTA, HSPA1A
43	ATP/ITP metabolism	2.044E-03	POLR1B, POLR1A, RPA39, RPB5, NDPK A, ADSS, HPRT, RPB7.0, 5'-NT1A, ADAR1, RRM1, RPB8
44	Transcription_CoREST complex-mediated epigenetic gene silencing	2.209E-03	BAF57, ZNF217, HDAC2, EHMT1, BAF170, c-Myb
45	DNA damage_ATM/ATR regulation of G2/M checkpoint: cytoplasmic signaling	2.609E-03	Chk2, Chk1, PARN, UBE2C, Cyclin B1, CDK1 (p34), 14-3-3
46	Signal transduction_mTORC2 upstream signaling	2.671E-03	TELO2, Rictor, HSP70, TTI1, GBL, EGFR, SIN1, Frizzled
47	Mitogenic action of Estradiol / ESR1 (nuclear) in breast cancer	3.337E-03	CAD, CHD8, NCOA3 (pCIP/SRC3), CDK2, Oct-1
48	Signal transduction_AKT(PKB) activation	3.551E-03	Rictor, Cyclin A, SETDB1, HSP90, GBL, EGFR, SIN1, CDK2
49	Cell cycle_Senescence activation pathways	3.628E-03	Chk2, Chk1, p14ARF, RHEB2, HSP90, p16INK4, CDK2
50	Aberrant B-Raf signaling in melanoma progression	4.027E-03	Rictor, B-Raf, RHEB2, HES1, MAP2, SRp55, CDK1 (p34)