## **Appendices**

Gene Name	Gene Description	Adjusted <i>p</i> - value (Benjamini– Hochberg)	Log 2 fold change ( <i>nol9<sup>sa1022</sup></i> /wild- type)
nol9	nucleolar protein 9	0.0127	-1.16
senp3b	SUMO1/sentrin/SMT3 specific peptidase 3b	1.13x10 <sup>-5</sup>	1.14
las11	LAS1-like (S. cerevisiae)	1	-0.166935452
senp3a	SUMO1/sentrin/SMT3 specific peptidase 3a	1	0.098424375
tex10	testis expressed 10	1	0.451582269
wdr18	WD repeat domain 18	1	0.357977686

**Table A - 1 The list of genes encoding proteins that interact with Nol9 in** *nol9*<sup>sa1022</sup> **mutants.** The gene name and description, the adjusted *p*-value and the fold change of mutant versus wild-type sibling are shown.

GO Term ID	Term	K-S value
GO:0048565	digestive tract development	0.0043
GO:0001889	liver development	0.02733
GO:0055123	digestive system development	0.04959
GO:0031017	exocrine pancreas development	0.11301
GO:0031016	pancreas development	0.14306
GO:0048546	digestive tract morphogenesis	0.29657
GO:0031018	endocrine pancreas development	0.47669

**Table A - 2 The list of GO terms associated with the digestive organs in** *nol9*<sup>sa1022</sup> **mutants.** The GO Term ID and name and the K-S value using topGO are shown.

Gene Name	Name Gene Description		Log 2 fold change ( <i>nol9<sup>sa1022</sup></i> /wild- type)
yars	tyrosyl-tRNA synthetase	1.38x10 <sup>-7</sup>	1.42
tars	threonyl-tRNA synthetase	5.78x10 <sup>-6</sup>	1.22
qars	glutaminyl-tRNA synthetase	5.87x10 <sup>-5</sup>	1.27
eprs	glutamyl-prolyl-tRNA synthetase	8.17x10 <sup>-5</sup>	1.17
iars	isoleucyl-tRNA synthetase	8.35x10 <sup>-5</sup>	1.28
kars	lysyl-tRNA synthetase	0.000389147	0.97
iars	isoleucyl-tRNA synthetase	0.000619223	1.48
hars	histidyl-tRNA synthetase	0.0007359	0.99
farsa	phenylalanyl-tRNA synthetase, alpha subunit	0.001796306	0.94
rars	arginyl-tRNA synthetase	0.005236793	0.89
farsb	phenylalanyl-tRNA synthetase, beta subunit	0.013378278	0.86
nars	asparaginyl-tRNA synthetase	0.028617189	0.81
larsb	leucyl-tRNA synthetase b	0.030845695	0.84

Table A - 3 The differentially expressed genes belonging to the KEGG term 'Aminoacyl-tRNA biosynthesis' in  $nol9^{a1022}$  mutants. The gene name and description, the adjusted *p*-value and the fold change of mutant versus wild-type sibling are shown.

Gene Name	Gene Description	Adjusted <i>p</i> - value (Benjamini– Hochberg)	Log 2 fold change ( <i>nol9<sup>sa1022</sup></i> /wild- type)
ctps1a	CTP synthase 1a	2.03x10 <sup>-8</sup>	1.57
POLR1D	polymerase (RNA) I polypeptide D, 16kDa	7.10x10 <sup>-7</sup>	1.56
cad	carbamoyl-phosphate synthetase 2, aspartate transcarbamylase, and dihydroorotase	2.77x10 <sup>-5</sup>	1.22
polr1c	polymerase (RNA) I polypeptide C	0.000232932	1.31
polr1e	polymerase (RNA) I polypeptide E	0.000331795	1.22
upp2	uridine phosphorylase 2	0.000487591	-1.37
POLR3D	polymerase (RNA) III (DNA directed) polypeptide D, 44kDa	0.00213729	1.09
znrd1	zinc ribbon domain containing 1	0.00557888	1.52
txnrd1	thioredoxin reductase 1	0.005973931	0.85
zgc:110540	zgc:110540	0.014289753	-0.87
polr1a	polymerase (RNA) I polypeptide A	0.036743601	0.90
umps	uridine monophosphate synthetase	0.044072318	1.05
znrd1	zinc ribbon domain containing 1	0.04613515	1.65

**Table A - 4 The differentially expressed genes belonging to the KEGG term 'Pyrimidine metabolism in** *nol9*<sup>sa1022</sup> **mutants..** The gene name and description, the adjusted *p*-value and the fold change of mutant versus wild-type sibling are shown.

Gene Name	Gene Description	Adjusted <i>p</i> - value (Benjamini– Hochberg)	Log 2 fold change ( <i>nol9<sup>sa1022</sup></i> /wild- type)
POLR1D	polymerase (RNA) I polypeptide D, 16kDa	7.10x10 <sup>-7</sup>	1.56
polr1c	polymerase (RNA) I polypeptide C	0.000232932	1.31
polr1e	polymerase (RNA) I polypeptide E	0.000331795	1.22
POLR3D	polymerase (RNA) III (DNA directed) polypeptide D, 44kDa	0.00213729	1.09
znrd1	zinc ribbon domain containing 1	0.00557888	1.52
polr1a	polymerase (RNA) I polypeptide A	0.036743601	0.90
znrd1	zinc ribbon domain containing 1	0.04613515	1.65

 Table A - 5 The differentially expressed genes belonging to the KEGG term 'RNA polymerase' in nol9<sup>sa1022</sup> mutants.

 The gene name and description, the adjusted p-value and the fold change of mutant versus wild-type sibling are shown.

Gene Name	Gene Description	Adjusted <i>p</i> - value (Benjamini– Hochberg)	Log 2 fold change ( <i>nol9<sup>sa1022</sup></i> /wild- type)
las11	LAS1-like (S. cerevisiae)	1	-0.448606006
nol9	nucleolar protein 9	1	0.229556284
senp3a	SUMO1/sentrin/SMT3 specific peptidase 3a	1	0.236577597
senp3b	SUMO1/sentrin/SMT3 specific peptidase 3b	1	0.273886684
tex10	testis expressed 10	1	0.070232461
wdr18	WD repeat domain 18	1	0.618400968

**Table A - 6 The list of genes encoding proteins that interact with Las11 in** *las11*<sup>sa674</sup> **mutants.** The gene name and description, the adjusted *p*-value and the fold change of mutant versus wild-type sibling are shown.

GO Term ID	Term	K-S value
GO:0031016	pancreas development	0.00447
GO:0048565	digestive tract development	0.00941
GO:0031017	exocrine pancreas development	0.0102
GO:0001889	liver development	0.0472
GO:0048546	digestive tract morphogenesis	0.13953
GO:0055123	digestive system development	0.25312
GO:0031018	endocrine pancreas development	0.28337

Table A - 7 The list of GO terms associated with the digestive organs in *las11<sup>sa674</sup>* mutants. The GO Term ID and name and the K-S value using topGO are shown.

KEGG Term		es	Fold Enrichment	n-value	Adjusted <i>p</i> -value
		%		p-value	Hochberg)
Aminoacyl-tRNA biosynthesis	16	2.46	7.14	2.64x10 <sup>-10</sup>	2.77 x10 <sup>-8</sup>
PPAR signaling pathway	11	1.69	3.72	4.08x10 <sup>-4</sup>	0.021191437
Steroid biosynthesis	6	0.92	6.82	9.34x10 <sup>-4</sup>	0.032179842
Pyrimidine metabolism	12	1.84	2.42	0.008207939	0.194544645
Glutathione metabolism	7	1.08	3.50	0.01145079	0.21482917
Purine metabolism	14	2.15	1.99	0.019360768	0.289748562
Steroid hormone biosynthesis	5	0.77	4.46	0.020533482	0.267438789
Drug metabolism		0.92	3.57	0.021400618	0.247182337
Spliceosome		2.30	1.86	0.025918624	0.263887035
RNA polymerase	5	0.77	3.92	0.032950845	0.296587998

**Table A - 8 The enriched KEGG terms of differentially expressed genes of** *tti*<sup>s450</sup> **mutants at a statistical significance of** *p***-value less than 0.05 using a modified Fisher's exact test.** The KEGG term, the number of genes involved in the terms and the percentage of involved genes over total number of genes, the fold enrichment, the *p*-value and the adjusted *p*-value using Benjamini-Hochberg procedure are shown.

KEGG Term		es	Fold Enrichment	<i>p</i> -value	Adjusted <i>p</i> -value (Benjamini-
		%			Hochberg)
Aminoacyl-tRNA biosynthesis	15	2.81	7.87	3.45x10 <sup>-10</sup>	3.38x10 <sup>-8</sup>
PPAR signaling pathway	11	2.06	4.58	6.45x10 <sup>-5</sup>	0.003154475
Steroid biosynthesis	7	1.31	8.26	6.99x10 <sup>-5</sup>	0.002281903
Glutathione metabolism		1.31	3.96	0.006238015	0.142136284
Retinol metabolism		1.13	4.05	0.012889366	0.22452057
Steroid hormone biosynthesis	5	0.94	5.06	0.013361798	0.197251552
Purine metabolism	13	2.44	2.09	0.017599578	0.220098841
RNA polymerase	5	0.94	4.16	0.026950046	0.284424572
p53 signaling pathway		1.50	2.52	0.034401259	0.316949433
Metabolism of xenobiotics by					
cytochrome P450	4	0.75	4.72	0.046546298	0.373189632

Table A - 9 The enriched KEGG terms of differentially expressed genes of  $set^{s453}$  mutants at a statistical significance of *p*-value less than 0.05 using a modified Fisher's exact test. The KEGG term, the number of genes involved in the terms and the percentage of involved genes over total number of genes, the fold enrichment, the *p*-value and the adjusted *p*-value using Benjamini-Hochberg procedure are shown.

GO Term ID	Term	K-S value
Digestive orga		
GO:0031017	exocrine pancreas development	0.04942
GO:0048546	digestive tract morphogenesis	0.05708
GO:0001889	liver development	0.10248
GO:0031018	endocrine pancreas development	0.22476
GO:0055123	digestive system development	0.30988
GO:0048565	digestive tract development	0.31777
GO:0031016	pancreas development	0.36157
Eye		
GO:0031076	embryonic camera-type eye development	0.19883
GO:0048596	embryonic camera-type eye morphogenesis	0.34133
GO:0048048	embryonic eye morphogenesis	0.64118
GO:0001754	eye photoreceptor cell differentiation	0.70683
GO:0048593	camera-type eye morphogenesis	0.72337
GO:0048592	eye morphogenesis	0.78997
GO:0042462	eye photoreceptor cell development	0.92979
GO:0043010	camera-type eye development	0.95017
GO:0001654	eye development	0.95104
GO:0060041	retina development in camera-type eye	0.96029
GO:0060042	retina morphogenesis in camera-type eye	0.98179
Brain		
GO:0030902	hindbrain development	0.3064
GO:0007420	brain development	0.6693
GO:0030900	forebrain development	0.87958
GO:0030901	midbrain development	0.893
Cartilage		
GO:0051216	cartilage development	0.12518
Skeletal		
GO:0001501	skeletal system development	0.0204
		0.26733
GO:0048706	embryonic skeletal system development	
GO:0060538	skeletal muscle organ development	0.29114
		0.29273
GO:0048704	embryonic skeletal system morphogenesis	
		0.41744
GO:0048705	skeletal system morphogenesis	
GO:0007519	skeletal muscle tissue development	0.48451
GO:0048741	skeletal muscle fiber development	0 75617

 Table A - 10 The list of GO terms associated with the digestive organs, eye, brain, cartilage and skeletal muscle in tti<sup>s450</sup> mutants. The GO Term ID and name and the K-S value using topGO are shown.

GO Term ID	Term	K-S value			
Digestive orga	Digestive organs				
GO:0048565	digestive tract development	0.02479			
GO:0055123	digestive system development	0.02639			
GO:0001889	liver development	0.17341			
GO:0031016	pancreas development	0.19378			
GO:0031017	exocrine pancreas development	0.27094			
GO:0048546	digestive tract morphogenesis	0.28365			
GO:0031018	endocrine pancreas development	0.31304			
Eye					
GO:0001654	eye development	0.08402			
GO:0042462	eye photoreceptor cell development	0.17352			
GO:0048048	embryonic eye morphogenesis	0.18294			
GO:0043010	camera-type eye development	0.22899			
GO:0048592	eye morphogenesis	0.24795			
GO:0001754	eye photoreceptor cell differentiation	0.26204			
GO:0031076	embryonic camera-type eye development	0.46281			
GO:0048596	embryonic camera-type eye morphogenesis	0.55498			
GO:0048593	camera-type eye morphogenesis	0.64427			
GO:0060041	retina development in camera-type eye	0.75822			
GO:0060042	retina morphogenesis in camera-type eye	0.89389			
Cartilage					
GO:0051216	cartilage development	0.27115			
Skeletal					
GO:0007519	skeletal muscle tissue development	0.00688			
GO:0001501	skeletal system development	0.03697			
GO:0048741	skeletal muscle fiber development	0.04216			
GO:0060538	skeletal muscle organ development	0.0603			
GO:0048705	skeletal system morphogenesis	0.12181			
GO:0048704	embryonic skeletal system morphogenesis	0.14073			
GO:0048706	embryonic skeletal system development	0.17811			

 Table A - 11 The list of GO terms associated with the digestive organs, eye, brain, cartilage and skeletal muscle in set<sup>s453</sup> mutants. The GO Term ID and name and the K-S value using topGO are shown.

Gene Name	Gene Description	Adjusted <i>p</i> - value (Benjamini–	Log 2 fold change ( <i>nol9<sup>sa1022</sup></i> /wild-			
150		Hochberg)	type)			
<i>tti<sup>s450</sup></i> mut	ant	1	1			
atg16l1	ATG16 autophagy related 16-like 1 (S. cerevisiae)	0.388405483	0.746690463			
atg4b	ATG4 autophagy related 4 homolog B (S. cerevisiae)	0.488785713	0.817575752			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	0.611158802	-0.772274754			
atg5	ATG5 autophagy related 5 homolog (S. cerevisiae)	0.788039883	0.513492925			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	0.818836971	-0.470201462			
atg4c	autophagy-related 4C (yeast)	0.876251962	-0.476068028			
atg10	ATG10 autophagy related 10 homolog (S. cerevisiae)	0.982750495	0.153372409			
atg13	ATG13 autophagy related 13 homolog (S. cerevisiae)	0.983675464	-0.049780343			
<i>set<sup>s453</sup></i> mu	itant		•			
atg16l1	ATG16 autophagy related 16-like 1 (S. cerevisiae)	0.609960781	0.648996789			
atg10	ATG10 autophagy related 10 homolog (S. cerevisiae)	0.81602828	0.940493174			
atg5	ATG5 autophagy related 5 homolog (S. cerevisiae)	0.833919644	0.638757882			
atg12	ATG12 autophagy related 12 homolog (S. cerevisiae)	0.994165769	0.652995621			
atg4b	ATG4 autophagy related 4 homolog B (S. cerevisiae)	0.99897654	0.516784842			
atg13	ATG13 autophagy related 13 homolog (S. cerevisiae)	0.999674861	0.226912478			
atg4c	autophagy-related 4C (yeast)	0.999674861	-0.204736068			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	0.999674861	0.217419046			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	0.999674861	-0.630197512			
nol9 <sup>sa1022</sup>	mutant					
atg4b	ATG4 autophagy related 4 homolog B (S. cerevisiae)	1	-0.18992073			
atg13	ATG13 autophagy related 13 homolog (S. cerevisiae)	1	0.190786591			
atg16l1	ATG16 autophagy related 16-like 1 (S. cerevisiae)	1	0.249922058			
atg10	ATG10 autophagy related 10 homolog (S. cerevisiae)	1	0.486924121			
atg4c	autophagy-related 4C (yeast)	1	0.241821933			
atg5	ATG5 autophagy related 5 homolog (S. cerevisiae)	1	0.174871634			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	1	-0.212406566			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	1	-0.095397584			
las11 <sup>sa674</sup>	las11 <sup>sa674</sup> mutant					
atg4b	ATG4 autophagy related 4 homolog B (S. cerevisiae)	1	0.014990544			
atg13	ATG13 autophagy related 13 homolog (S. cerevisiae)	1	0.137029624			
atg16l1	ATG16 autophagy related 16-like 1 (S. cerevisiae)	1	0.112731025			
atg4c	autophagy-related 4C (yeast)	1	0.599221648			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	1	-0.081416492			
atg10	ATG10 autophagy related 10 homolog (S. cerevisiae)	1	0.162001033			
atg5	ATG5 autophagy related 5 homolog (S. cerevisiae)	1	0.152707837			
atg9a	ATG9 autophagy related 9 homolog A (S. cerevisiae)	1	-0.755265462			

**Table A - 12 The list of** *autophagy-related genes (atg)* in *nolo<sup>sa1022</sup>, las11<sup>sa674</sup>, ttis<sup>s450</sup>* and *set<sup>453</sup>* mutants. The gene name and description, the adjusted *p*-value and the fold change of mutant over wild-type sibling are shown.



## **Enriched Cellular Component Ontology Terms**

Figure A - 1 The enriched cellular component ontology terms in *tti<sup>s450</sup>* mutants at statistical significance of K-S value less than 0.01 (-log10 (K-S value) more than 2.0).





-logio (K-S value)



## **Enriched Biological Process Ontology Terms**

Figure A - 3 The enriched biological process ontology terms in tts<sup>3450</sup> mutants at statistical significance of K-S value less than 0.01 (-log10 (K-S value) more than 2.0).



Figure A - 4 The enriched cellular component ontology terms in  $set^{s453}$  mutants at statistical significance of K-S value less than 0.01 (-log10 (K-S value) more than 2.0).



**Enriched Molecular Function Ontology Terms** 



Figure A - 6 The enriched biological process ontology terms in set<sup>s453</sup> mutants at statistical significance of K-S value less than 0.01 (-log10 (K-S value) more than 2.0