

A

<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>
<i>his3Δ est1Δ</i>	<i>est1Δ est1Δ</i>	<i>rad57Δ est1Δ</i>	<i>est3Δ est1Δ</i>	<i>tel1Δ est1Δ</i>	<i>rad55Δ est1Δ</i>	<i>rad54Δ est1Δ</i>	<i>asf1Δ est1Δ</i>	<i>xrs2Δ est1Δ</i>	<i>dcc1Δ est1Δ</i>	<i>rif1Δ est1Δ</i>	<i>his3Δ est1Δ</i>
<i>his3Δ est1Δ</i>	<i>rif2Δ est1Δ</i>	<i>elg1Δ est1Δ</i>	<i>pol32Δ est1Δ</i>	<i>upf2Δ est1Δ</i>	<i>upf3Δ est1Δ</i>	<i>cdd73Δ est1Δ</i>	<i>ydl118Δ est1Δ</i>	<i>sum1Δ est1Δ</i>	<i>rfm1Δ est1Δ</i>	<i>rtf1Δ est1Δ</i>	<i>his3Δ est1Δ</i>
<i>his3Δ est1Δ</i>	<i>hst3Δ est1Δ</i>	<i>spt21Δ est1Δ</i>	<i>hmo1Δ est1Δ</i>	<i>mot3Δ est1Δ</i>	<i>upf1Δ est1Δ</i>	<i>kem1Δ est1Δ</i>	<i>lea1Δ est1Δ</i>	<i>mak10Δ est1Δ</i>	<i>mak31Δ est1Δ</i>	<i>mak3Δ est1Δ</i>	<i>his3Δ est1Δ</i>
<i>his3Δ est1Δ</i>	<i>rrp8Δ est1Δ</i>	<i>mrpl44Δ est1Δ</i>	<i>csm3Δ est1Δ</i>	<i>sla1Δ est1Δ</i>	<i>rad27Δ est1Δ</i>	<i>chk1Δ est1Δ</i>	<i>ebs1Δ est1Δ</i>	<i>rad9Δ est1Δ</i>	<i>rad24Δ est1Δ</i>	<i>rad52Δ est1Δ</i>	<i>his3Δ est1Δ</i>
<i>his3Δ est1Δ</i>	<i>mre11Δ est1Δ</i>	<i>exo1Δ est1Δ</i>	<i>rad17Δ est1Δ</i>	<i>hex3Δ est1Δ</i>	<i>bnr1Δ est1Δ</i>	<i>rho4Δ est1Δ</i>	<i>ede1Δ est1Δ</i>	<i>bmh1Δ est1Δ</i>	<i>bmh2Δ est1Δ</i>	<i>ddc1Δ est1Δ</i>	<i>his3Δ est1Δ</i>
<i>his3Δ est1Δ</i>	<i>ctf18Δ est1Δ</i>	<i>ctf8Δ est1Δ</i>	<i>slx8Δ est1Δ</i>	<i>tsa1Δ est1Δ</i>	<i>sgs1Δ est1Δ</i>	<i>rtt107Δ est1Δ</i>	<i>rtt101Δ est1Δ</i>	<i>mms1Δ est1Δ</i>	<i>mms22Δ est1Δ</i>	<i>rtt109Δ est1Δ</i>	<i>his3Δ est1Δ</i>
<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>	<i>his3Δ est1Δ</i>

Red: Telomere length (Askree *et al.* 2004; Gatbonton *et al.* 2006);

Purple: DNA replication or repair (Collins *et al.* 2008);

Green: Genes of general interest.

B Passage1

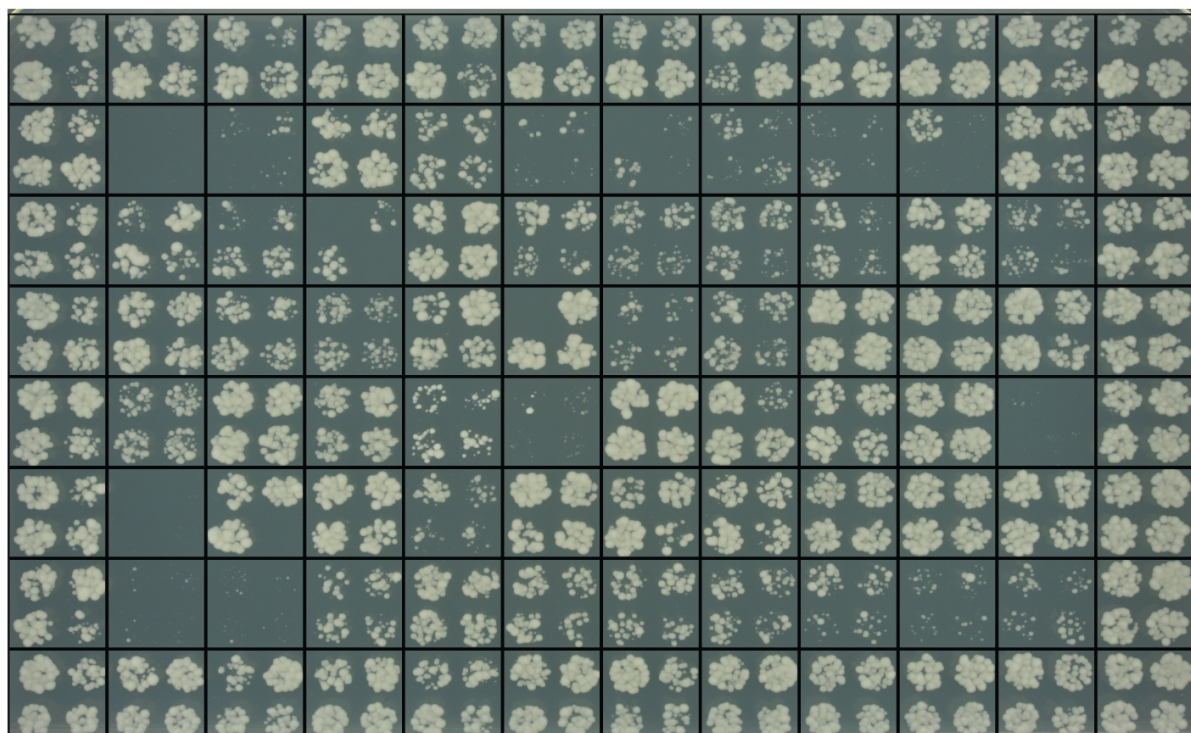


Figure S1 60 *yfgΔ est1Δ* strains arrayed in quadruplicate on a 384 format plate. (A) Strain map indicating the position of 60 genotypes on the 384 format plate in panel B; four replicates drawn in light grey. Red, purple, and green names indicate the reason for choosing particular genes for this study. (B) Photograph of a 384 format plate at passage 1 (48 hours after inoculation) from the liquid procedure (Figure 1A)

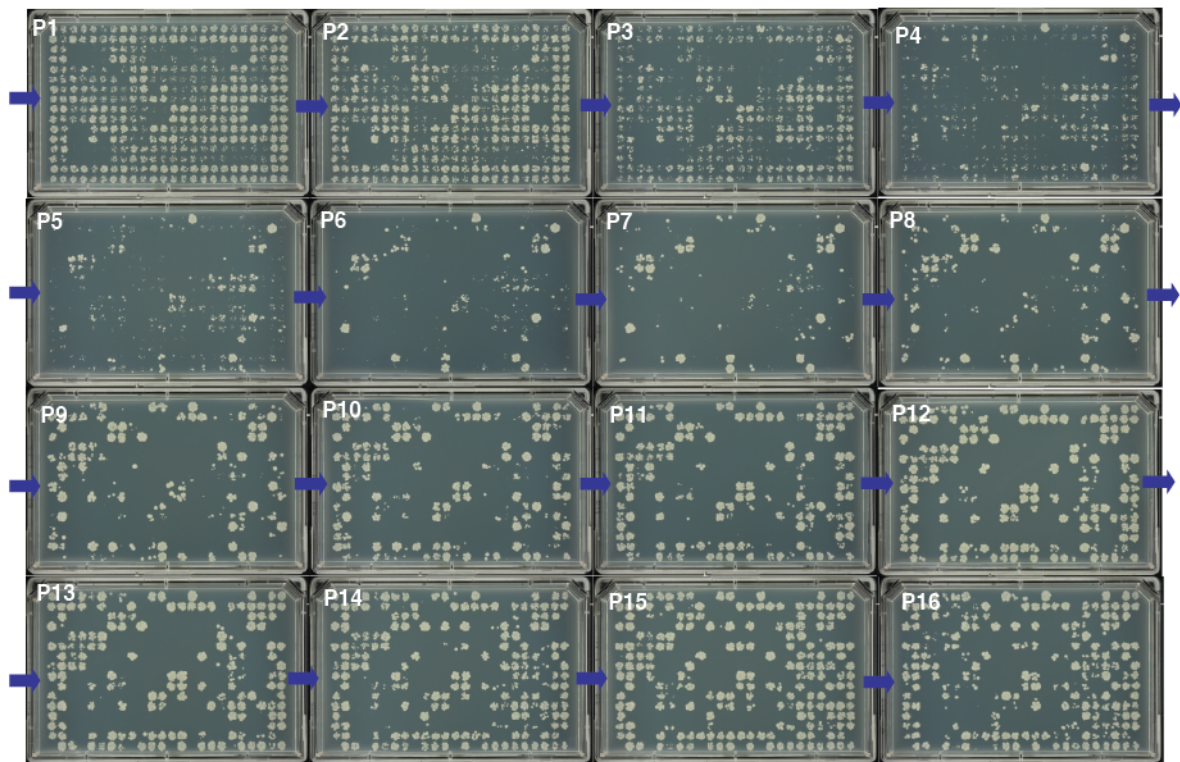
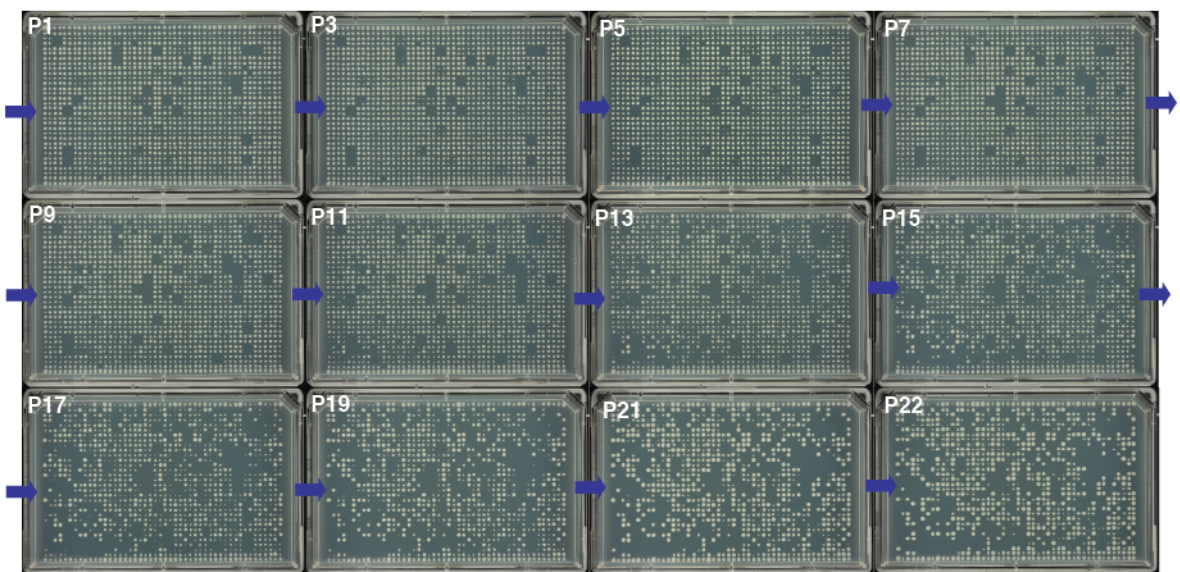
A**B**

Figure S2 Example photographs from passaged cultures. (A) Photographs of 384 format agar plates for each of the 16 passages from the liquid procedure. (B) 12 sample photographs at various passages (indicated) for one of the 1536 format plates from the solid procedure. Passage numbers are labelled on each photograph; P1-passage 1, etc.

A

	P1	P2	P3	P4	P5	P6	P7	P8	P9	P10	P11	P12	P13
WT	314	330	334	344	330	384	460	442	436	372	286	610	400
<i>est3Δ</i>	369	240	137	15	7	29	510	282	362	482	292	378	399
<i>est1Δ</i>	410	269	143	23	9	44	521	301	332	387	297	400	365
<i>est3Δ est1Δ</i>	212	120	85	19	76	110	310	230	305	386	255	388	303

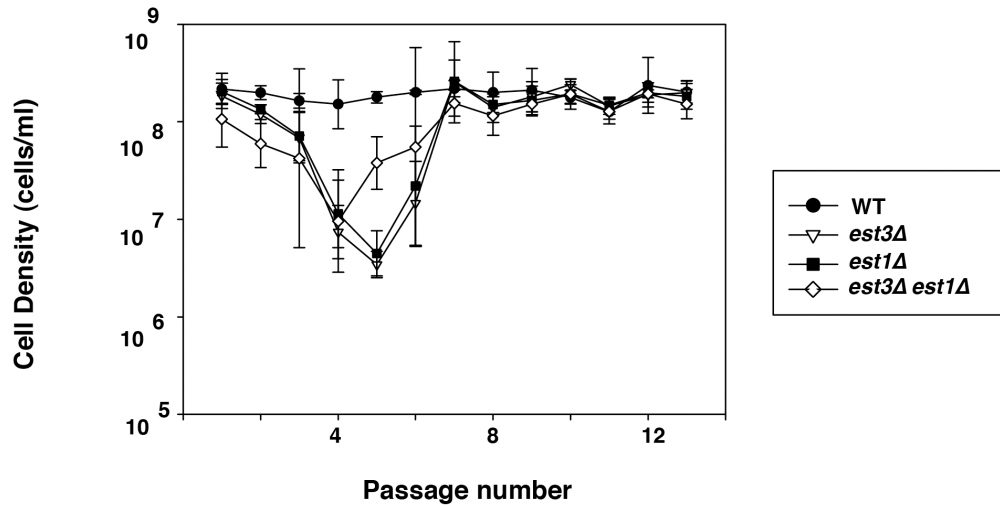
B

Figure S3 Low through-put senescence experiment in the W303 genetic background. (A) Six independent strains of each genotype in the W303 background were taken directly from germination plates and grown in liquid culture. Cell densities were counted every 23 hrs, followed by dilution to 5×10^5 cells/ml and continued incubation. Cell density reached by cells with the same genotype after 23 hours were recorded and averaged. The dilution factors at each passage were calculated and showed in the table, genotypes are indicated. (B) The data from (A) were plotted. Each symbol represents the average cell density reached by cells with the same genotype after 23 hours and the error bars represent the standard deviations.

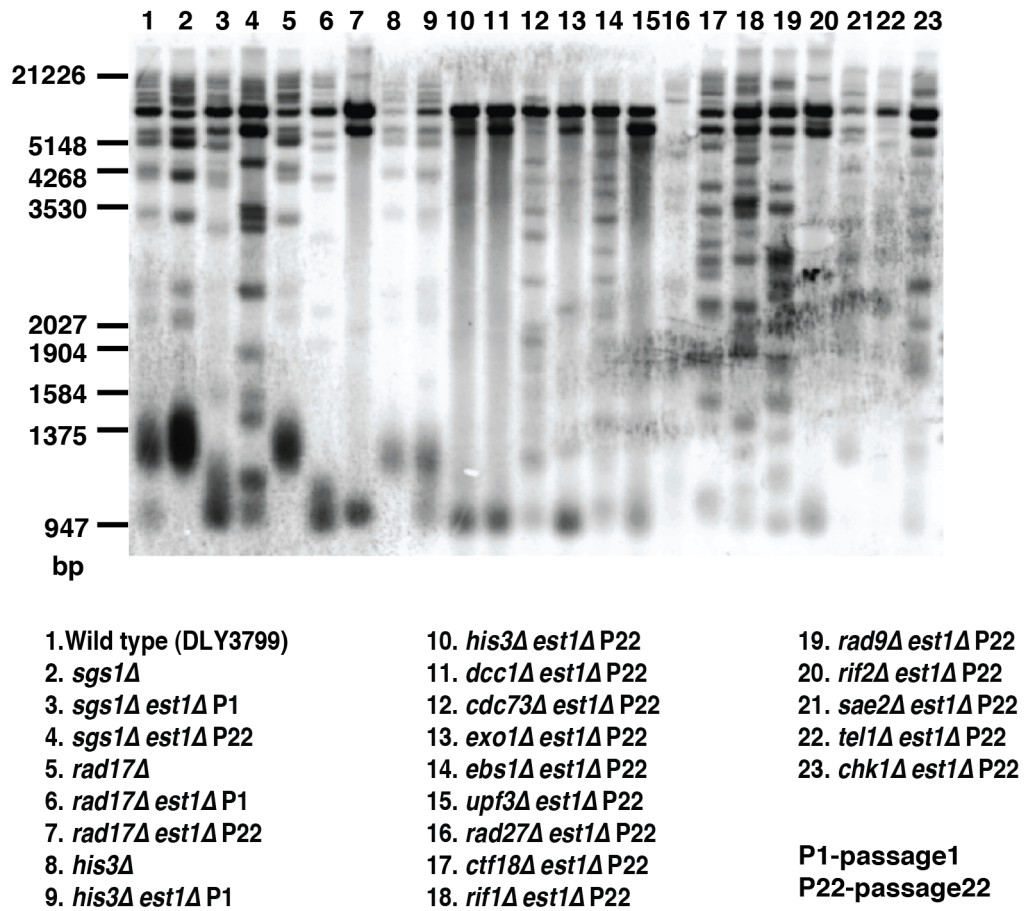
A

Figure S4 Survivors were produced by passage 22 in the solid procedure. Genomic DNA prepared from the strains of different genotypes as indicated, digested with *Xho*I and subjected to Southern blot to detect telomeric 'Y' and 'TG' fragments.

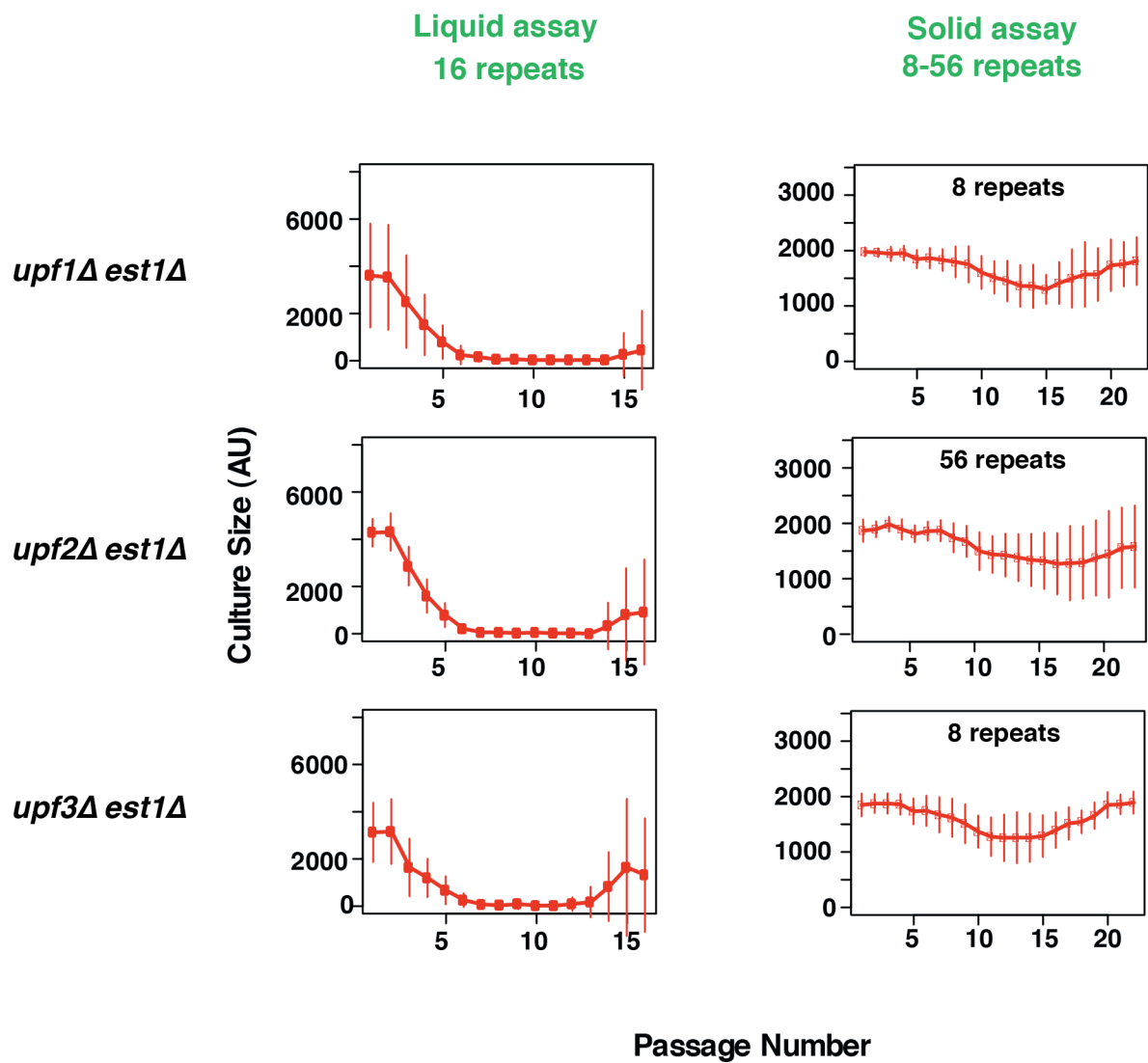


Figure S5 Nonsense mediated decay genes had consistent MDPs in both screens. *UPF1*, *UPF2*, and *UPF3* gene deletions had consistent MDPs patterns in both the liquid procedure and in the solid procedure. Replicate number and experiment type are indicated in green text.

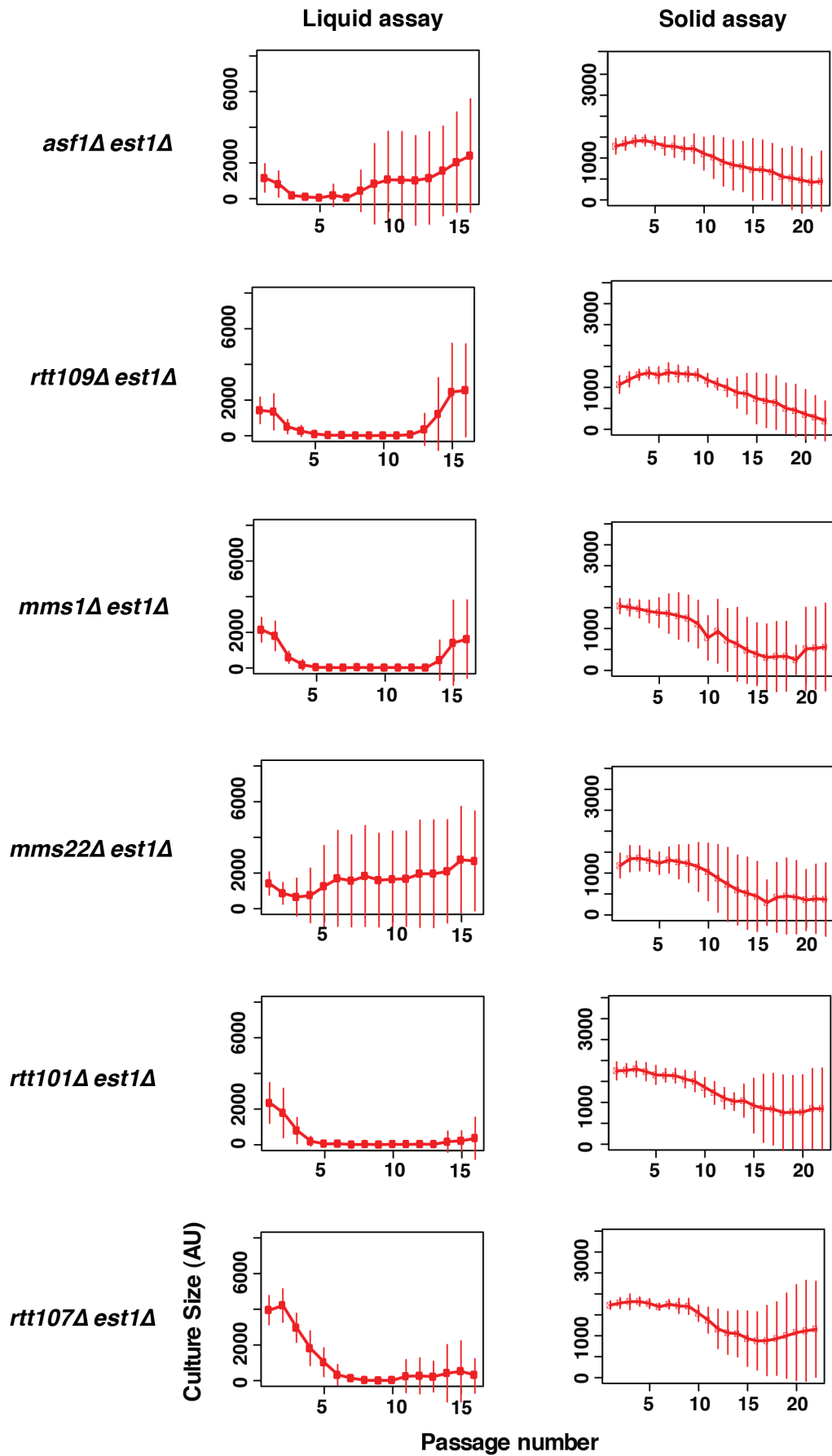


Figure S6 Genes affecting replication fork progression had an accelerated senescence phenotype when deleted in the *est1Δ* background. MDPs for *asf1Δ est1Δ*, *rtt109Δ est1Δ*, *mms1Δ est1Δ*, *mms22Δ est1Δ*, *rtt101Δ est1Δ*, and *rtt107Δ est1Δ* indicating a fast senescence phenotype in both the liquid and solid assay.

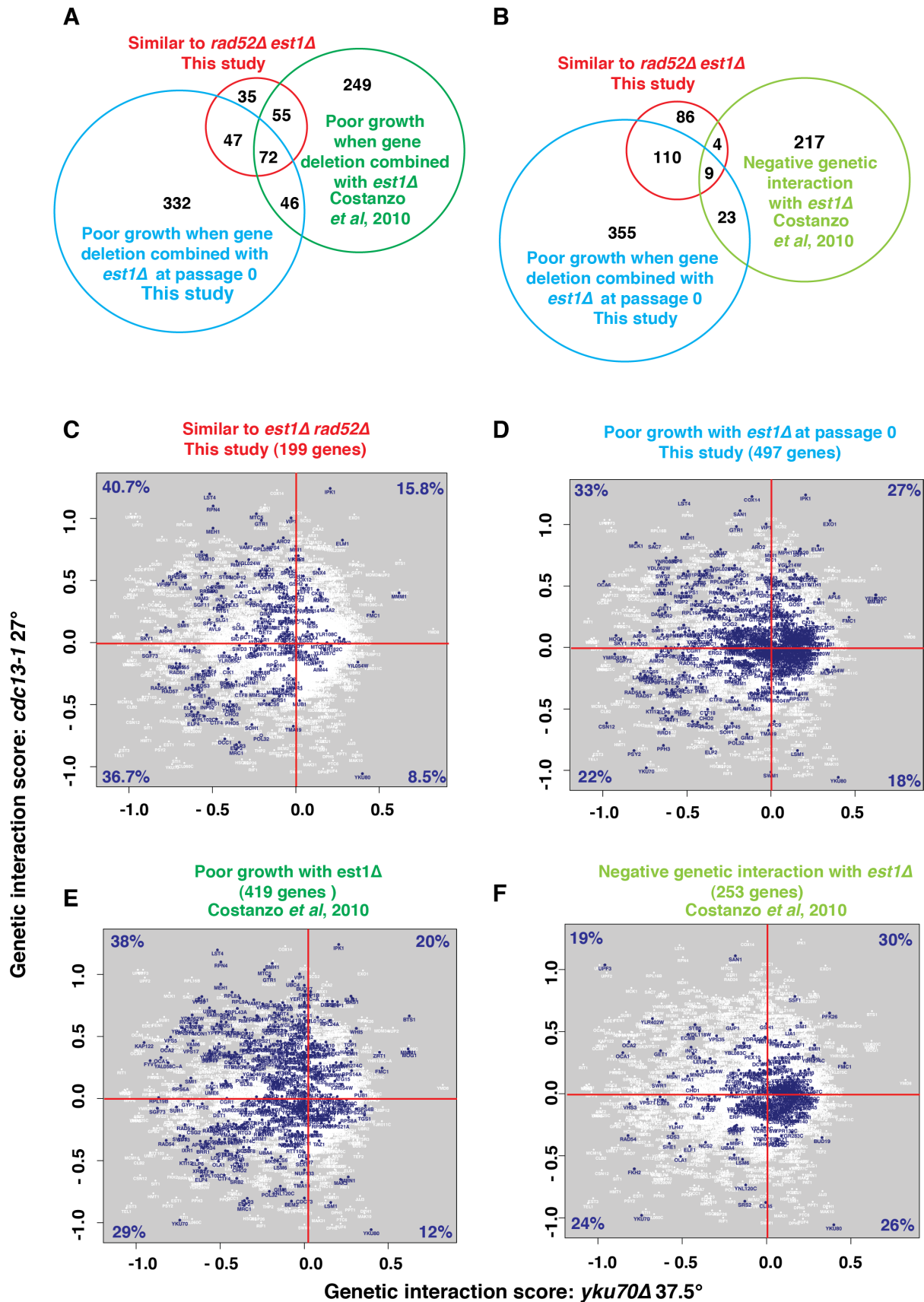


Figure S7 Repeated passage of cultures provides useful information about telomere-dependent senescence. (A) Venn diagram demonstrating the overlap between poor growers from our initial SGA (blue circle), poor growers from the *est1Δ* SGA by (COSTANZO *et al.* 2010b) (green circle) and our accelerated senescence class defined by repeated passaging, similar to *rad52Δ*, Figure 4B, Supporting File S7 (red circle). Poor growers are defined here as deletions whose fitness is more than one standard deviation less than the mean fitness across all deletions in the library. (B) As for panel A,

except the green circle in this panel represents genes classified as having a lenient negative interaction with *est1Δ* by Costanzo et al. (C) *yku70Δ* vs *cdc13-1* genetic interaction profile (see Figure 6) with our accelerated senescence class overlaid (blue). (D) *yku70Δ* vs *cdc13-1* genetic interaction profile with poor growers from our initial SGA overlaid in blue (E) *yku70Δ* vs *cdc13-1* genetic interaction profile with poor growers from the *est1Δ* SGA by Costanzo et al. overlaid (blue) (F) *yku70Δ* vs *cdc13-1* genetic interaction profile with genes classified as having a lenient negative interaction with *est1Δ* by Costanzo et al. overlaid (blue, see Figure 4A, Table ST4).

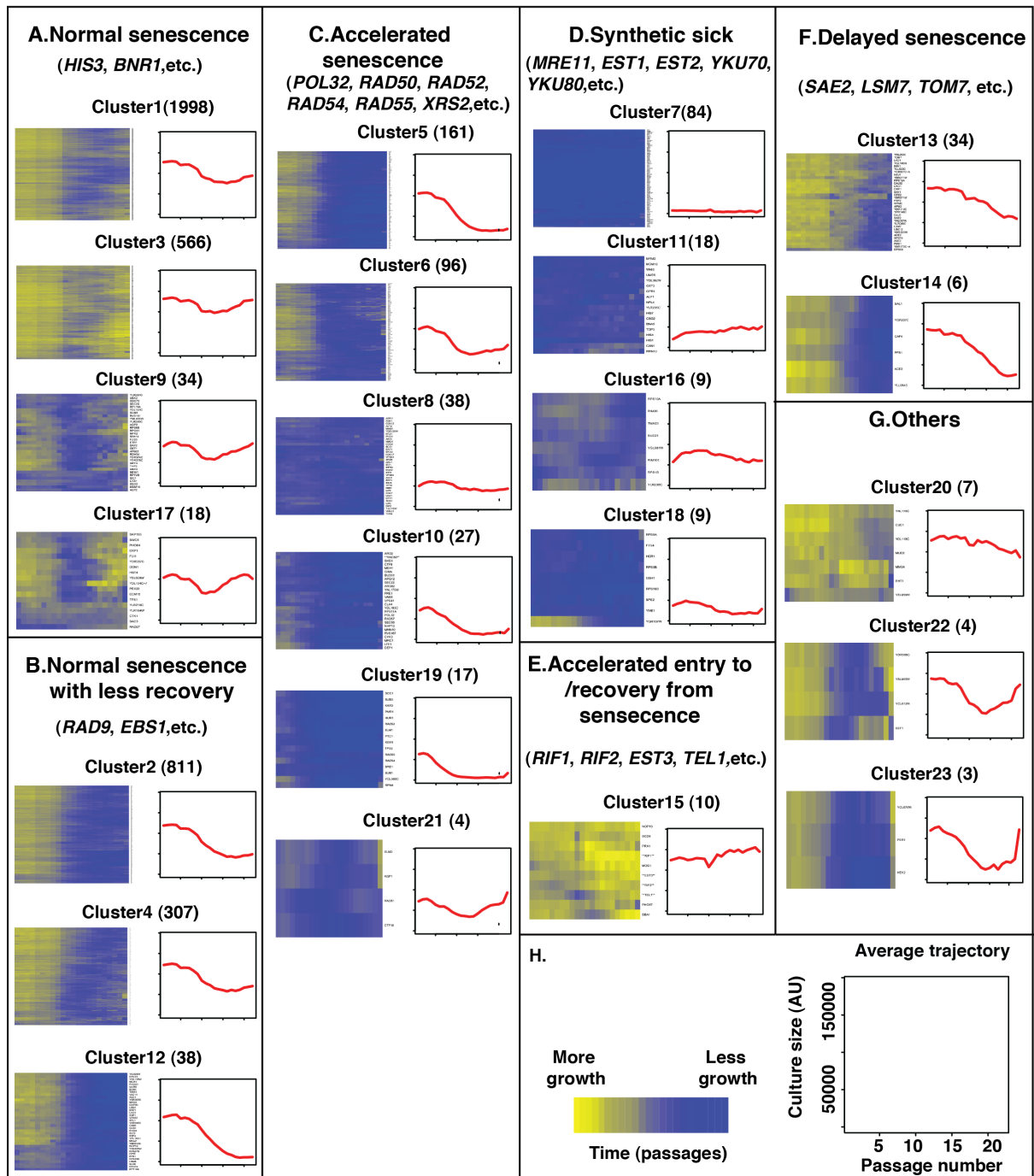


Figure S8 Unsupervised hierarchical QT clustering identifies 23 different MDP classes. Heat map representation of cluster MDPs on left, average MDPs on right. 23 automatically generated classes were combined manually into 8 functionally meaningful classifications (A-H).

Files S1-S13

All supporting files listed below (together with raw images and initial culture size quantifications) can be downloaded from our Supplemental Data website: <http://research.ncl.ac.uk/colonyzer/ChangSenescence/>

File S1. Liquid screen MDP plots.

Mean Density Profiles for each of 60 strains grown in the liquid culture screen. Red points are average culture size for all available repeats at passages 1 - 16. Vertical red lines show standard deviation for all repeats.

File S2. Solid screen MDP plots.

Mean Density Profiles for each of ~4300 strains grown in the solid agar screen. Red points are average culture size for all available repeats at passages 1 - 22. Vertical red lines show standard deviation for all repeats.

File S3. Manual classification of MDPs from liquid screen.

Manual functional clustering of the 60 gene deletions investigated in the liquid culture screen. MDPs for each cluster member plotted in panels A – P. 3 main categories are indicated in black text; functionally related sub-categories are defined by panels and titled in green text: (A) Recombination repair (B) *CTF18* RFC-like complex (C) Nonsense mediated decay. (D) Histone acetyltransferase. (E) MRX complex. (F) *SLX5-SLX8* STUbL Complex. (G) Ubiquitin-conjugating enzyme. (H) AF1 complex. (I) *SUM1/RFM1* repressor complex. (J) Mean plots of gene deletions that accelerate senescence. (K) Mean plots of gene deletions that accelerated entry to and escaped from senescence. (L) Subunit of N-terminal acetyltransferase. (M) DNA damage checkpoint. (N) 14-3-3 proteins. (O&P) Gene deletions that do not affect senescence. Dashed lines indicate genes that are not classified in any particular group.

File S4. MDPs for all 4299 strains in the solid screen.

Tab-delimited text file containing quantitative MDPs (culture size profiles with increasing passage). Culture sizes are in arbitrary units. Note that photo 1 corresponds to the first photograph taken after the SGA. Passage 1 in the manuscript corresponds to photo 2.

File S5. MDPs similar to that of the *his3Δ est1Δ* archetype from the solid screen.

Tab-delimited text file listing Pearson's correlation coefficient (*c*) between the MDP of the archetype strain *his3Δ est1Δ* and all other deletions in the library. Gene deletions are ranked by correlation coefficient and thereby profile similarity. Root Mean Square difference (RMS) between the MDP of each strain and that of the archetype is also reported, together with an estimate of the area under the MDP of each strain (Integral).

File S6. MDPs similar to that of the *rad52Δ est1Δ* archetype from the solid screen.

As for File S5.

File S7. MDPs similar to that of the *rad52Δ est1Δ* archetype from the solid screen which are also poor growers.

As for S5, except similarity is defined as deletions in the library with $\text{Corr} > 0.5$ and Culture Size at passage 2 < 1680 (see Figure 4).

File S8. MDPs similar to that of the *rif1Δ est1Δ* archetype from the solid screen.

As for File S5.

File S9. Gene lists used in the Venn diagram in Figure S7

Tab-delimited lists of gene names classified as having similar MDPs to the *rad52Δ est1Δ* archetype (RAD52Archetype), classified as being poor growers after the SGA presented in this work (SGASick), classified as being poor growers in the *est1Δ* double mutant fitness data set Costanzo et al. 2010 (Supporting data file S1) (CostanzoSick) or classified as having a (lenient) negative interaction with *est1Δ* by Costanzo et al. 2010 (CostanzoNegGIS). Poor growers were defined as having a fitness or culture size less than the mean by more than one standard deviation.

File S10. Intersection classes in Venn diagram Figure S7.

Venn diagram from Figure S7 repeated with letters classifying groups of gene deletions from overlapping gene lists as defined in File S9.

File S11. Gene members of interaction classes in Venn diagram in File S10.

Tab-delimited member lists for gene deletion categories defined in File S10.

File S12. 23 clusters from unsupervised QT clustering of MDPs from solid screen.

Clusters are represented by MDP heat maps showing culture size variation with passage (V1-V22), dendograms linking MDPs (left) the names of gene deletions for each MDP (right) together with alternating plots of the average MDP for all members of each cluster (red curve, passage number on x-axis, culture size on y-axis).

File S13. Gene members of clusters presented in File S12

Tab-delimited text file classifying each ORF in the deletion library by cluster membership.

Table S1 60 genes tested by liquid senescence assay

Gene	*Senescence phenotype	**Telomere length screens	Gene ontology annotation	***References
<i>EST3</i>	AERS	VS	telomerase component	<i>est2Δ est3Δ/est1Δ est3Δ/tlc1Δ est3Δ</i> showed normal senescence (LENDVAY <i>et al.</i> 1996)
<i>RIF1</i>	AERS	VL	telomere binding protein	<i>rif1Δ est2Δ</i> showed normal senescence (ANBALAGAN S <i>et al.</i> 2011); <i>rif1Δ est2Δ</i> accelerated entry to /recovery from senescence (CHANG <i>et al.</i> 2011)
<i>RIF2</i>	AERS	VL	telomere binding protein	<i>rif2Δ est2Δ</i> accelerated entry to/recovery from senescence (CHANG <i>et al.</i> 2011)
<i>EST1</i>	AS	VS	telomerase component	
<i>MRE11</i>	AS	VS	DNA repair/ MRX complex	Normal (LE <i>et al.</i> 1999)
<i>XRS2</i>	AS	VS	DNA repair/MRX complex	Normal (LE <i>et al.</i> 1999)
<i>RAD52</i>	AS		recombinational repair of double-strand breaks in DNA	<i>rad52Δ est1Δ and rad52Δ tlc1Δ</i> accelerated senescence (ABDALLAH <i>et al.</i> 2010; LE <i>et al.</i> 1999; LEBEL <i>et al.</i> 2009; LEE <i>et al.</i> 2007; LUNDBLAD and BLACKBURN 1993)
<i>RAD54</i>	AS		recombinational repair of double-strand breaks in DNA	<i>rad54Δ tlc1Δ</i> has severe growth defect (LE <i>et al.</i> 1999)
<i>RAD55</i>	AS		recombinational repair of double-strand breaks in DNA	
<i>RAD57</i>	AS		recombinational repair of double-strand breaks in DNA	<i>rad57Δ tlc1Δ</i> has severe growth defect (LE <i>et al.</i> 1999)
<i>UPF1 (NAM7)</i>	AS	VS	nonsense mediated decay	<i>upf1Δ</i> with <i>tlc1Δ/est1Δ/est2Δ/est3Δ</i> delayed senescence measured by a different method (ENOMOTO <i>et al.</i> 2004)
<i>UPF2 (NMD2)</i>	AS	S	nonsense mediated decay	<i>upf2Δ</i> with <i>tlc1Δ/est1Δ/est2Δ/est3Δ</i> delayed senescence measured by a different method (ENOMOTO <i>et al.</i> 2004)
<i>UPF3</i>	AS	S	nonsense mediated decay	<i>upf3Δ</i> with <i>tlc1Δ/est1Δ/est2Δ/est3Δ</i> delayed senescence measured by a different method (ENOMOTO <i>et al.</i> 2004)

<i>DCC1</i>	AS	ss	Sister chromatid cohesion	
<i>CTF18</i>	AS	Literature reported	Sister chromatid cohesion	
<i>CTF8</i>	AS	ss	Sister chromatid cohesion	
<i>ELG1</i>	AS	L	RFC complex	
<i>RTT101</i>	AS		Histone acetyltransferase/involved in NHEJ	
<i>RTT109</i>	AS		Histone acetyltransferase/involved in NHEJ	
<i>RTT107</i>	AS		Mms22-dependent DNA repair during S phase/interacts with Mms22p and Slx4p	
<i>MMS1</i>	AS		Subunit of an E3 ubiquitin ligase complex/resolving replication intermediates	<i>mms1Δ tlc1Δ</i> accelerated senescence and failed to recover (ABDALLAH <i>et al.</i> 2010)
<i>MMS22</i>	AS		Ubiquitin-conjugating enzyme variant involved in error-free post replication repair	
<i>CDC73</i>	AS	S	PAF1 complex	
<i>RTF1</i>	AS	ss	PAF1 complex	
<i>SLX8</i>	AS	sl,	Subunit of the Slx5-Slx8 SUMO-targeted ubiquitin ligase (STUbL) complex	<i>slx8Δ tlc1Δ</i> accelerated senescence (AZAM <i>et al.</i> 2006)
<i>SLX5 (HEX3)</i>	AS		Subunit of the Slx5-Slx8 SUMO-targeted ubiquitin ligase (STUbL) complex	<i>slx5Δ tlc1Δ</i> accelerated senescence (AZAM <i>et al.</i> 2006)

<i>RRP8</i>	AS	L	pre-rRNA processing/methyltransferase	
<i>SPT21</i>	AS	ss	regulator of histone gene transcription	
<i>POL32</i>	AS	sl	Polymerase delta subunit	<i>pol32Δ tlc1Δ</i> accelerated senescence (LYDEARD <i>et al.</i> 2007)
<i>SGS1</i>	AS		Nucleolar DNA helicase of the RecQ family	<i>sgs1Δ tlc1Δ</i> accelerated senescence (AZAM <i>et al.</i> 2006; LEE <i>et al.</i> 2007)
<i>RAD27</i>	AS	VL	flap-endonuclease	<i>rad27Δ</i> with <i>est1Δ/tlc1Δ/est3Δ/cdc13-2</i> accelerated senescence (PARENTEAU and WELLINGER 2002)
<i>HMO1</i>	AS	L	HMG-box protein	
<i>TEL1</i>	AS	VS	PIK homologue	<i>tel1Δ tlc1Δ</i> showed normal senescence (ENOMOTO <i>et al.</i> 2002) ; <i>tel1Δ tlc1Δ</i> delayed senescence (RITCHIE <i>et al.</i> 1999), <i>tel1Δ tlc1Δ mec1Δ sml1Δ</i> is normal (CHAN <i>et al.</i> 2001); <i>tel1Δ tlc1Δ</i> delayed senescence (ABDALLAH <i>et al.</i> 2010)
<i>KEM1</i>	AS	S	RNA degradation	
<i>YDL118</i>	AS	ss	Unknown	
<i>LEA1</i>	AS	L	RNA splicing	
<i>SUM1</i>	AS	S	SUM1/RFM1 repressor complex	
<i>RFM1</i>	AS	S	SUM1/RFM1 repressor complex	
<i>TSA1</i>	AS		Thioredoxin peroxidase/ribosome-associated and free cytoplasmic antioxidant	
<i>ASF1</i>	AS		Nucleosome assembly factor/chromatin assembly and disassembly	
<i>SLA1</i>	AS		Cytoskeletal binding protein	

<i>RHO4</i>	AS		Non-essential small GTPase of the Rho/Rac subfamily of Ras-like proteins	
<i>EDE1</i>	AS		Key endocytic protein	
<i>EXO1</i>	Normal	Literature reported	5'-3' exonuclease and flap-endonuclease	<i>exo1Δ tlc1Δ</i> delayed senescence (MARINGELE and LYDALL 2004)
<i>EBS1</i>	Normal	ss	nonsense mediated decay	
<i>RAD9</i>	Normal	Literature reported	DNA damage checkpoint effector	<i>rad9Δ tlc1Δ</i> had less G2/M arrested cells (IJPMA and GREIDER 2003)
<i>RAD17</i>	Normal	Literature reported	DNA damage checkpoint effector	
<i>DDC1</i>	Normal	Literature reported	DNA damage checkpoint effector	
<i>RAD24</i>	Normal	Literature reported	DNA damage checkpoint effector	<i>rad24Δ tlc1Δ</i> had less G2/M arrested cells (IJPMA and GREIDER 2003)
<i>CHK1</i>	Normal		DNA damage checkpoint effector	
<i>BMH1</i>	Normal	Literature reported	14-3-3 protein	
<i>BMH2</i>	Normal	Literature reported	14-3-3 protein	
<i>MRPL44</i>	Normal	ss	mitochondrial ribosomal protein	
<i>MOT3</i>	Normal	ss	POLII transcription	
<i>HST3</i>	Normal		Member of the Sir2 family of NAD(+)-dependent protein deacetylases	
<i>CSM3</i>	Normal		Replication fork associated factor	
<i>BNR1</i>	Normal		Formin	
<i>HIS3</i>	Normal		Imidazoleglycerol-phosphate dehydratase	Control strain
<i>MAK3</i>	Normal	L	N-terminal acetyltransferase complex	
<i>MAK10</i>	Normal	L	N-terminal acetyltransferase complex	
<i>MAK31</i>	Normal	L	N-terminal acetyltransferase complex	

Legend: Examining of the 60 genes selected for the liquid procedure in this study. The references quoted were indicated in the table and included below.

*Senescence phenotype abbreviations: AS- accelerated senescence, AERS-accelerated entry to and recovery from senescence.

Normal: normal entry to and recovery from senescence.

** Telomere phenotype abbreviations (ASKREE *et al.* 2004; GATBONTON *et al.* 2006; SHACHAR *et al.* 2008): VS - very short, S - short, ss - slightly short, sl - slightly long, L - long, VL - very long.

*** Conflicts in data may be due to differences in ways senescence was measured.

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Table S2 Gene ontology analysis (processes) of RAD52-like genes from the solid screen in this study

GO-Slim term	Genes annotated to the term
Transport	<i>AGP1, APQ12, ATG9, AVL9, AVT5, CCZ1, CIK1, CLA4, DRS2, FKS1, GCS1, GET2, GOS1, GTR1, KAR3, LST4, MMM1, NUM1, PEX12, PEX14, PEX19, PEX5, PMR1, RIM8, RPS18B, SEC28, SEC66, SGF73, SKY1, SLG1, SNX4, SPC2, SPF1, SSO2, STE20, THR4, TIM18, TLG2, UBP3, UBX2, VAM3, VAM6, VAM7, VPS1, VPS4, VPS41, VPS5, VPS51, VTH1, YDL183C, YPT7, ZRC1</i>
Response to stress	<i>ASF1, ATG9, BRE1, CAC2, CCZ1, CKB1, CTF18, CTF4, CTF8, DCC1, ELM1, GSH1, HDA1, MGA2, MMS1, MMS22, MRC1, MRE11, NHP10, NPL4, OPI1, PHO5, PHO80, POL32, RAD50, RAD52, RAD54, RAD55, RAD57, RMI1, RPN4, RTT109, SIC1, SIN3, SLG1, SLX5, SLX8, SNF6, SOH1, STB5, STE20, TAX4, TIM18, TMA19, TPS2, UBI4, URM1, VAM3, VPS75, XRS2, YKU80</i>
RNA metabolic process	<i>ASF1, BRE1, CAC2, CKB1, DEG1, ELP2, ELP3, ELP4, ELP6, GRS1, GTR1, HAP4, HDA1, HTZ1, INO2, IPK1, MED1, MGA2, MMS1, MRC1, MRE11, NCS6, NOP12, OPI1, PHO80, POL32, RPL35B, RPN4, RPS11B, RPS16A, RPS18B, RTF1, RTT109, SDS3, SGF11, SGF73, SIN3, SKY1, SNF4, SNF6, SOH1, STB5, SWD1, SWD3, URM1, YGR122W, YKU80, YPL205C, ZDS2</i>
Chromosome organization	<i>ARP6, ASF1, BRE1, CAC2, CIK1, CTF18, CTF4, CTF8, DCC1, HDA1, KAR3, LGE1, MMS22, MRC1, NHP10, RAD50, RAD51, RAD52, RAD54, RAD57, RMI1, RTF1, RTS1, RTT109, SDS3, SGF11, SGF29, SGF73, SIN3, SLX5, SLX8, SNF6, SOH1, SWD1, SWD3, VPS75, XRS2, YKU80</i>
Transcription, DNA-dependent	<i>ASF1, BRE1, CAC2, CKB1, ELP2, ELP3, ELP4, ELP6, GRS1, GTR1, HAP4, HDA1, HTZ1, INO2, MED1, MGA2, MRC1, MRE11, OPI1, PHO80, RPN4, RTF1, RTT109, SDS3, SGF11, SGF73, SIN3, SNF4, SNF6, SOH1, STB5, SWD1, SWD3, YGR122W, YKU80, YPL205C, ZDS2</i>
Protein modification process	<i>ASF1, BRE1, CKB1, CLA4, DIA2, ELM1, ELP2, ELP6, HDA1, HPM1, LGE1, MUB1, NCS6, OST4, PHO80, RTF1, RTS1, RTT109, SDS3, SGF11, SGF29, SGF73, SIC1, SIN3, SKY1, SLX5, SLX8, SNF4, STE20, SWD1, SWD3, UBI4, UBP3, UBP6, URM1, VPS75</i>
DNA metabolic process	<i>BRE1, CAC2, CTF18, CTF4, CTF8, DCC1, DIA2, HUR1, LGE1, MMS1, MMS22, MRC1, MRE11, NHP10, POL32, RAD50, RAD51, RAD52, RAD54, RAD55, RAD57, RIM1, RPN4, RTT109, SIN3, SLX5, SLX8, SNF6, SOH1, SWD1, SWD3, VPS75, XRS2, YKU80</i>
Cell cycle	<i>BRE1, CIK1, CLA4, CTF18, CTF4, CTF8, DCC1, ELM1, FAR1, GCS1, KAR3, MMS22, MRC1, MRE11, PHO80, RAD50, RAD51, RAD52, RAD55, RAD57, RIM8, RMI1, RPN4, RTS1, SAP155, SHE1, SIC1, SIN3, SMI1, SOH1, STE20, XRS2, ZDS2</i>
Vesicle-mediated transport	<i>AVL9, CCZ1, DRS2, FKS1, GCS1, GET2, GOS1, LST4, PMR1, RIM8, SEC28, SLG1, SNX4, SSO2, THR4, TLG2, UBP3, VAM3, VAM6, VAM7, VPS1, VPS4, VPS41, VPS5, VPS51, VTH1, YPT7</i>
Cellular membrane organization	<i>APQ12, ATG9, CCZ1, DRS2, FKS1, GET2, GOS1, MEH1, MMM1, RIM8, SEC28, SLG1, SNX4, SSO2, THR4, TIM18, TLG2, VAM10, VAM3, VAM6, VAM7, VPS1, VPS4, VPS41, YPT7</i>
Biological process unknown	<i>AIM4, AIM44, IES2, IES5, MTC1, MTC5, PAR32, RTC2, YBR174C, YDR149C, YGL024W, YGL081W, YGR125W, YGR182C, YIL054W, YLR065C, YLR108C, YLR287C, YML012C-A, YNL170W, YPL102C</i>
Meiosis	<i>BRE1, CIK1, KAR3, MMS22, MRE11, RAD50, RAD51, RAD52, RAD55, RAD57, RIM8, RTS1, SOH1, XRS2</i>

Response to chemical stimulus	<i>CLA4, ELM1, FAR1, GSH1, NPL4, OPI1, RPN4, SKY1, STB5, STE20, TIM18, TMA19, URM1</i>
Protein complex biogenesis	<i>ATG9, ATP11, COX12, COX23, FMC1, GIM4, PEX14, PEX5, PKR1, SGF73, VAM6, VPS4, VPS41</i>
Cellular amino acid metabolic process	<i>ARO1, ARO2, GRS1, GSH1, GSH2, HOM6, PDC5, SPE1, SPE2, SPE3, THR4, ZRC1</i>
Cellular lipid metabolic process	<i>APQ12, CHO2, ETR1, INO2, INP52, MGA2, OAR1, OPI1, OPI3, PSD1, SCT1, TLG2</i>
Peroxisome organization	<i>ATG9, CCZ1, PEX12, PEX14, PEX19, PEX5, SLG1, SNF4, SNX4, TLG2, VPS1, VPS51</i>
Chromosome segregation	<i>CIK1, CTF18, CTF4, CTF8, DCC1, KAR3, MMS22, MRC1, RMI1, RTS1, SOH1</i>
Mitochondrion organization	<i>ATP11, COX12, COX23, FMC1, GEM1, GRS1, MMM1, MRPL36, NUM1, RIM1, TIM18</i>
Translation	<i>EAP1, GRS1, MRPL36, RPL21B, RPL35B, RPS11B, RPS16A, RPS18B, RPS4A, TEF4, TMA19</i>
Vesicle organization	<i>GOS1, SEC28, SSO2, TLG2, VAM3, VAM6, VAM7, VPS4, VPS41, VPS51</i>
Vacuole organization	<i>ATG9, CLA4, STE20, VAM10, VAM3, VAM6, VAM7, VPS41, YPT7</i>
Cellular carbohydrate metabolic process	<i>ELM1, FKS1, MAL12, OST4, PDC5, PFK2, SNF4, SOL4, TPS2</i>
Signaling	<i>CLA4, FAR1, MRC1, OPI1, PHO80, SLG1, STE20, TAX4</i>
Sporulation resulting in formation of a cellular spore	<i>MRE11, RIM21, RIM9, SSO2, UBI4, XRS2, YPL205C</i>
Cytoskeleton organization	<i>CIK1, GCS1, NUM1, SHE1, SLG1, VPS1</i>
Ribosome biogenesis	<i>DRS2, NOP12, RPL35B, RPS11B, RPS16A, RPS18B</i>
Cytokinesis	<i>CLA4, CTS1, CYK3, ELM1, STE20</i>
Generation of precursor metabolites and energy	<i>CYT1, ETR1, OAR1, PDC5, PFK2</i>
Cofactor metabolic process	<i>CAT5, SOL4, SPE1, SPE2, SPE3</i>
Cell budding	<i>CLA4, ELM1, MUB1, URM1, VPS51</i>
Cellular homeostasis	<i>MEH1, PHO80, SKY1, SPF1, ZRC1</i>
Conjugation	<i>CIK1, FAR1, KAR3, STE20</i>
Pseudohyphal growth	<i>DFG16, ELM1, SOK2, STE20</i>
Fungal-type cell wall organization	<i>ECM8, KRE1, SLG1, TAX4</i>
Heterocycle metabolic process	<i>AAH1, IMD3, PDC5, SOL4</i>
Cellular aromatic compound metabolic process	<i>AAH1, ARO1, ARO2, PDC5</i>
Cellular protein catabolic process	<i>DIA2, NPL4, UBX2, VMS1</i>
Cellular respiration	<i>CYT1, ETR1, OAR1</i>
Nucleus organization	<i>APQ12, CIK1, KAR3</i>
Vitamin metabolic process	<i>SPE1, SPE2, SPE3</i>
Cellular component morphogenesis	<i>ELM1, SSO2</i>
Transposition	<i>MMS1, RTT109</i>
Protein folding	<i>EGD2</i>

Table legend: *RAD52*-like genes in this study were analysed using SGD gene ontology slim mapper

(<http://www.yeastgenome.org/cgi-bin/GO/goSlimMapper.pl>). “Yeast GO-Slim: Process” GO set was used to determine the list of genes that involved in different biological processes.