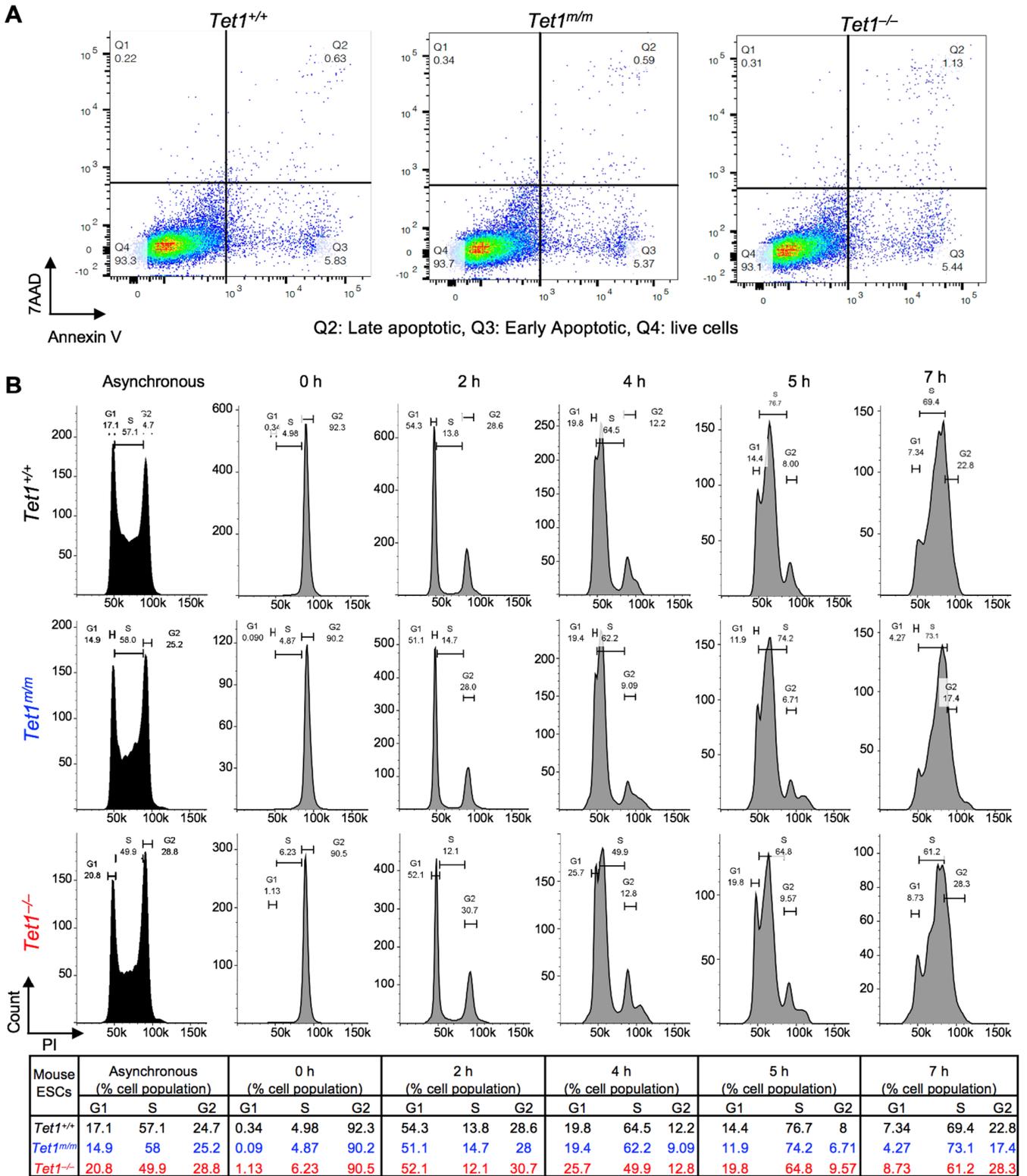
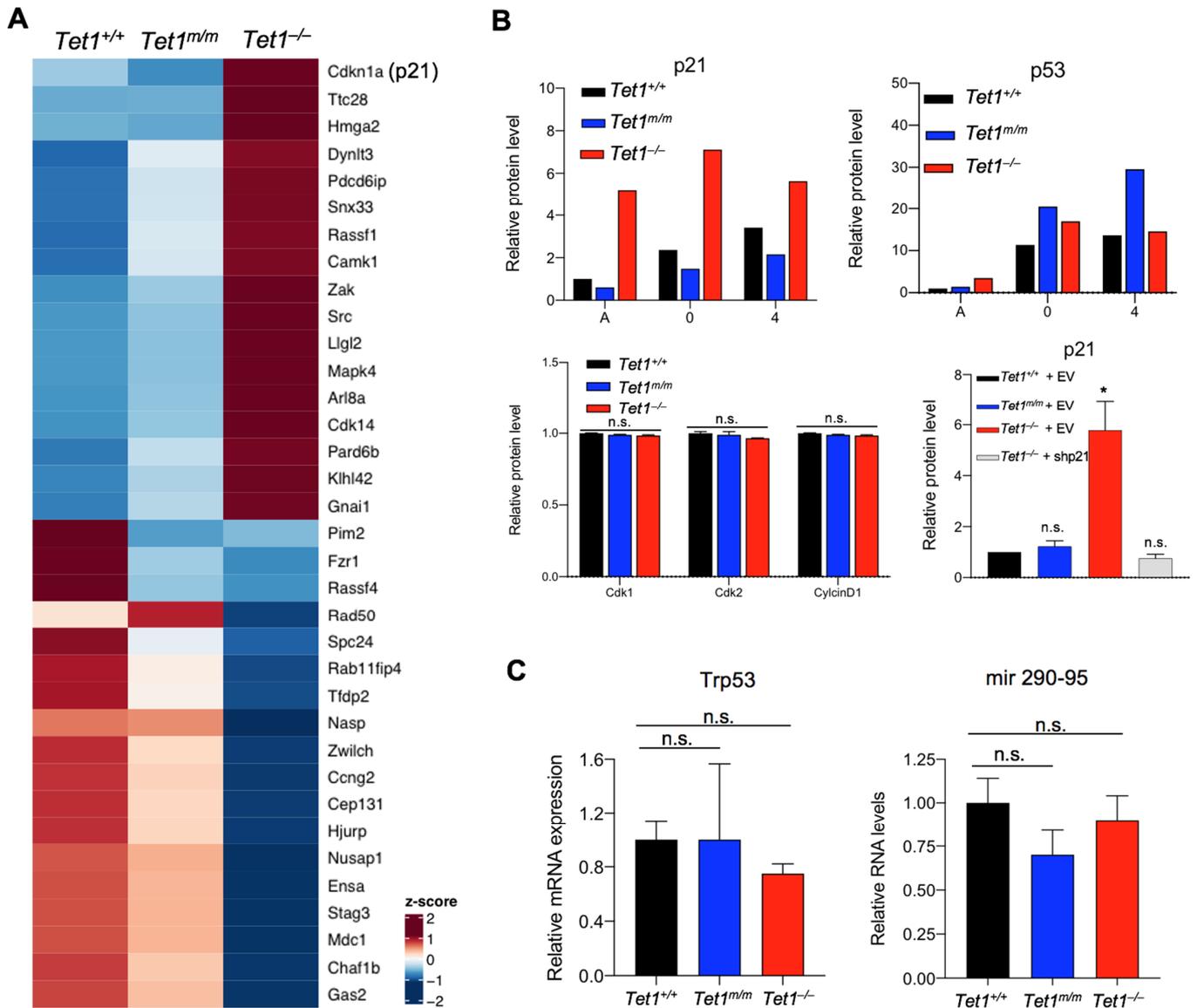


# Supplementary Information



**Figure S1:** Apoptosis and cell cycle profiles of *Tet1*<sup>+/+</sup>, *Tet1*<sup>m/m</sup> and *Tet1*<sup>-/-</sup> mESCs are analyzed by flow cytometry. (A) Analysis of Annexin V and 7AAD-stained ESCs of indicated genotypes by flow

cytometry; % cells in each quadrant are shown. (B) Cell cycle analysis of asynchronous ESCs and synchronized (Nocodazole block for 16 h) ESCs at indicated time-points after Nocodazole release. Cells were stained by PI and subjected to a flow cytometry analysis; % cells in each phase are summarized in Table S1 below. Note that in *Tet1*<sup>-/-</sup> ESCs there was reduced % ESCs in the S phase and increased % ESCs in the G1 phase, particularly at 4 and 5 h post-release.



**Figure S2:** mRNA and protein levels of cell cycle regulators are quantified in *Tet1*<sup>+/+</sup>, *Tet1*<sup>m/m</sup> and *Tet1*<sup>-/-</sup> mESCs. (A) Heatmap of the expression of genes in cell cycle regulation gene ontology terms, uniquely deregulated in *Tet1*<sup>-/-</sup>, but not in *Tet1*<sup>m/m</sup> and *Tet1*<sup>+/+</sup> ESCs based on our published RNA-seq data (see Methods). (B) Quantification of protein levels of cell cycle regulators as assessed by Western blot in Figures 2B, 2C and 2E. All signals were normalized with the corresponding actin signals. (C) Quantification of RNA levels of p53 and pri-mir290-95 cluster in ESCs of indicated genotypes by RT-

qPCR. Data normalized to *Gapdh*.  $n = 3$  ESC lines of each genotype. For all panels, error bars represent Stdev. \*: statistically significant; n.s.: not significant when compared with wild type or control.

**Table S1:** List of oligos used in study.

Name	Sequence (5'-3')	Purpose	Reference
Tet1 RT-qPCR For	TGCACCTACTGCAAGAATCG	Real time qPCR	[14]
Tet1 RT-qPCR Rev	AAATTGGCATCACAGCTTCC	Real time qPCR	[14]
Nanog RT-qPCR For	AAGCAGAAGATGCGGACTGT	Real time qPCR	[15]
Nanog RT-qPCR Rev	ATCTGCTGGGAGGCTGAGGTA	Real time qPCR	[15]
Pou5f1 RT-qPCR For	ACATCGCCAATCAGCTTGG	Real time qPCR	[15]
Pou5f1 RT-qPCR Rev	AGAACCATACTCGAACCACATCC	Real time qPCR	[15]
Bin1 RT-qPCR For	CAAGGCAAACACTACAGGCTCATC	Real time qPCR	[16]
Bin1 RT-qPCR Rev	CCACGTTTCATCTCCTCGAAC	Real time qPCR	[16]
Eomes RT-qPCR For	TGCAAGAGAAAGCGCCTGTCTC	Real time qPCR	[17]
Eomes RT-qPCR Rev	CAATCCAGCACCTTGAACGACC	Real time qPCR	[17]
Gapdh RT-qPCR For	GTGTTCTACCCCAATGTGT	Real time qPCR	[14]
Gapdh RT-qPCR Rev	ATTGTCATACCAGGAAATGAGCTT	Real time qPCR	[14]
p21 RT-qPCR For	CCATGAGCGCATCGCAATC	Real time qPCR	[18]
p21 RT-qPCR Rev	CCTGGTGATGTCCGACCTG	Real time qPCR	[18]
p27 RT-qPCR For	GGCCCGTCAATCATGAA	Real time qPCR	[19]
p27 RT-qPCR Rev	TTGCGTGACTCG CTTCTTC	Real time qPCR	[19]
p15 RT-qPCR For	GGTGGGTGCAGTCAGTACCT	Real time qPCR	[19]
p15 RT-qPCR Rev	CGAGCTGGAGGTGACTTCTC	Real time qPCR	[19]
p16 RT-qPCR For	CAACGCCCCGAACTCTTTC	Real time qPCR	[19]
p16 RT-qPCR Rev	GCAGAAGAGCTGCTACGTGAAC	Real time qPCR	[19]
p19 RT-qPCR For	CGGTATCCACTATGCTTCTGGAA	Real time qPCR	[19]
p19 RT-qPCR Rev	CCGCTGCGCCACTCAA	Real time qPCR	[19]
Trp53 RT-qPCR For	TGCTCACCTGGCTAAAGTT	Real time qPCR	[20]
Trp53 RT-qPCR Rev	GTCCATGCAGTGAGGTGATG	Real time qPCR	[20]
Pri-mir290-95 RT-qPCR For	GAACCTCACGGGAAGTGACC	Real time qPCR	[21]
Pri-mir290-95 RT-qPCR Rev	TGCCCACAGGAGAGACTCAA	Real time qPCR	[21]
p21 For	GGCCGACGCTATAAGGAGG	Tet1 ChIP-qPCR	This paper
p21 Rev	CTATTGTTCCCTGCCACGAAG	Tet1 ChIP-qPCR	This paper
p21 For	GTGTAGGAAGGTGACCAGGC	Ezh2/H3K27me3 ChIP-qPCR	This paper
p21 Rev	CCCCAAATGCCAAACCCAAG	Ezh2/H3K27me3 ChIP-qPCR	This paper