

Supplementary Information

RNA content in the nucleolus alters p53 acetylation *via* MYBBP1A

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Supplementary Methods

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Supplementary Methods

siRNAs

All siRNAs were purchased from Invitrogen. The siRNA duplexes siMYBBP (5'-UUCACCAGCCGACCUGACUGAAAGA-3'), siMYBBP-2 (5'-GGUCCGAGAUGAAAUAUGCCCUGAA-3'), siTIF-IA (5'-CGACACCGUGGUUUCUCAUGCCAAU-3'), siRPL5 (5'-CCUACUUUAAGAGAUACCAAGUGAA-3'), siRPL11 (5'-GCUAGAUACACUGUCAGAUCUUUG-3'), siRPL23 (5'-CAGCAGUGGUCAUUCGACAACGAAA-3'), siRPL26 (5'-AAUCCCUUUGUGACUUCGACCGAA-3'). Stealth™ RNAi Luciferase reporter control duplex was used as a control.

RT-qPCR primers

5'-ATCGTCCACCGCAAATGCTTCTA-3' and
5'-AGCCATGCCAATCTCATCTTGTT-3' for β -Actin,
5'-CCATGAAAAAGGACATAGTG-3' and 5'-CGTGTCAAAGGAGCTTGGT-3' for
TIF-IA, 5'-CCTGCAAACATGAGAAGCTG-3' and
5'-AGTGTGCCCCCTGCCTGTAG-3' for MYBBP1A,
5'-TGTTGCAGATTACATGCGCTACT-3' and
5'-CTCTCGTATAGCAGCATGAGCTT-3' for RPL5,
5'-CAGGTTTCAGCATCGCAGACAAG-3' and
5'-TTTGCCAGGAAGGATGATCCCA-3' for RPL11,
5'-AGTGGTCATTCGACAACGAAAG-3' and

5'-CTACTGGTCCTGTAATGGCAGAA-3' for RPL23,
5'-CGGGAAAAGGCTAATGGCACAAC-3' and
5'-CGAGATTTGGCTTTCCGTTTCGAG-3' for RPL26,
5'-GGAGACTCTCAGGGTCGAAA-3' and 5'-TTAGGGCTTCCTCTTGGAGA-3' for
p21, 5'-CTCTCAGATGAAGATGATGAGG-3' and
5'-CTGTTGCAATGTGATGGAAGG-3' for HDM2,
5'-GGGCCCAGACTGTGAATCCT-3' and
5'-ACGTGCTCTCTCTAAACCTATGCA-3' for PUMA, 5'-
AGTCGGGTTGCTTGGGAATGC -3' and 5'- CCCTTACGGTACTTGTTGACT -3'
for 28S rRNA.

Antibodies

Rabbit anti-human MYBBP1A antibody was raised against a synthetic peptide corresponding to 1265–1328 amino acids of human MYBBP1A. Anti-Myc (MC045) antibody was purchased from Nacalai tesque; anti-FLAG M2 antibody, anti-FLAG M2-agarose and anti-β-Actin antibody were from Sigma; anti-HA (3F10) and anti-BrdU antibodies were purchased from Roche diagnostics; anti-p53 (DO-1 and FL393), anti-HDM2 (SMP-14), anti-UBF (F-9) and anti-p300 (N-15) antibodies were from Santa Cruz Biotechnology; anti-p53-K382Ac, anti-p53-S15P and anti-PUMA antibodies were from Cell Signaling Technology; anti-p53-K305Ac antibody was from BioLegend; anti-p53-K373Ac antibody was from Epitomics; anti-NPM antibody was from Zymed Laboratories.

RNA purification and RT-qPCR

Total RNA was isolated from cultured cells with Sepasol RNA I Super reagent (Nacal Tesque) and subjected to reverse transcription using random hexamers and SuperScript III reverse transcriptase (Invitrogen) according to the manufacturer's protocol. RT-qPCR was performed with the Thermal Cycler Dice Real Time System (TaKaRa). Data analysis was performed by using the comparative Ct method. Results were normalized to β -actin.

Immunofluorescence

Cells grown on chamber slides were rinsed twice with phosphate-buffered saline (PBS) (140 mM NaCl, 2.7 mM KCl, 1.5 mM KH_2PO_4 , and 8.1 mM Na_2HPO_4) and fixed in 3.7% formaldehyde in PBS for 10 min. After rinsing twice with PBS, the cells were permeabilized in 0.1% Triton X-100 in PBS, blocked with TBS-T buffer containing 0.5% bovine serum albumin and 10% goat serum for 1 hour at room temperature. Then the cells were incubated with anti-MYBBP1A, anti-UBF, or anti-NPM antibodies for 1 hour, stained with AlexaFluor-conjugated secondary antibodies (Invitrogen) for 1 hour and mounted with Vectashield (Vector Laboratories). Immunofluorescence was performed using Biozero immunofluorescence microscopy (Keyence, Osaka, Japan).

Immunoblotting

Cell extracts were separated by SDS-polyacrylamide gel electrophoresis and transferred to PVDF membranes. After blocking with 3% skim milk in TBS-T buffer [20 mM Tris-HCl (pH 7.5), 150 mM NaCl, and 0.05% Tween20] for 1 hour, the membranes were incubated with the first antibody overnight at 4°C. After washing with TBS-T buffer, the membranes were incubated with horseradish peroxidase-conjugated

secondary antibody for 90 min. Bands were detected with Chemi-Lumi One (Nacalai tesque) or Immobilon Western blotting detection kit (Millipore). Stripping was performed using WB stripping solution (Nacalai tesque) according to the manufacturer's protocol. Total, phosphorylated, and acetylated forms of p53 were evaluated on the same membrane.

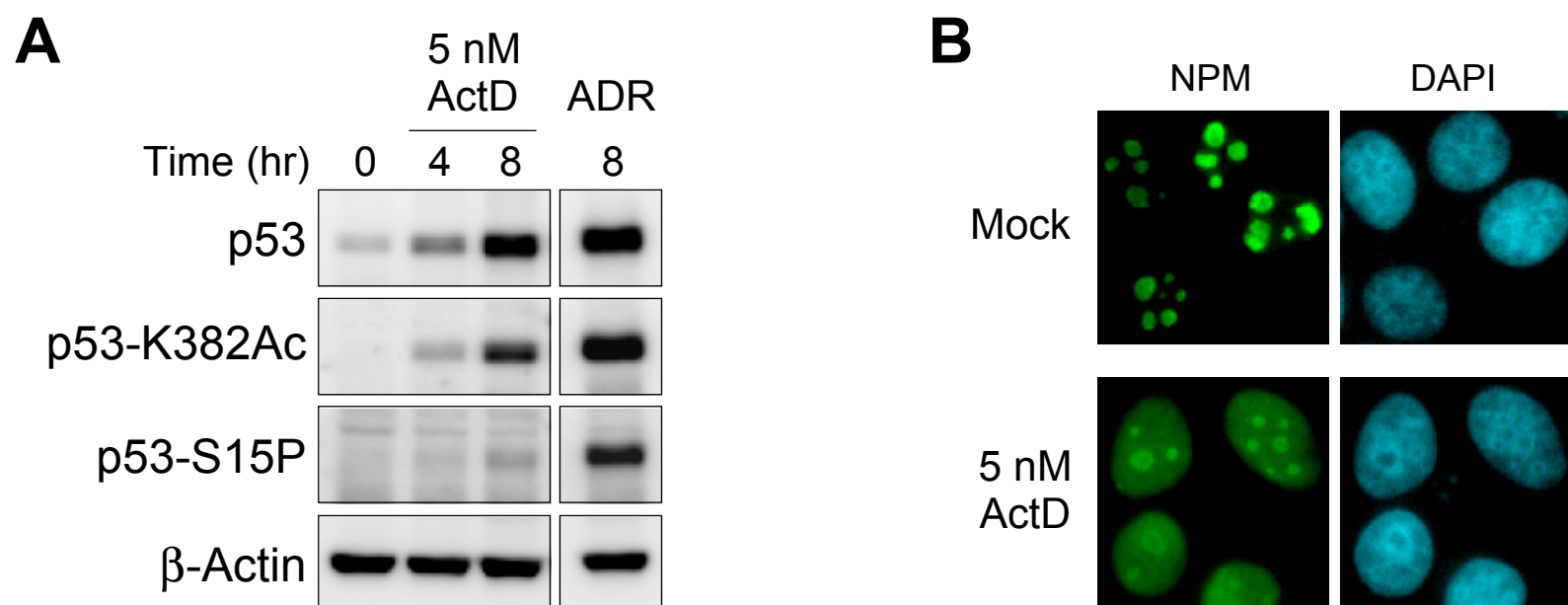
GST pull-down assay

cDNAs encoding full-length human p53 and its deletion mutant derivatives were cloned into pGEX-4T-1 (Amersham Biosciences). GST fusion proteins were expressed in BL-21 cells following induction with IPTG and purified with glutathione Sepharose 4B beads (Amersham). ³⁵S-labeled MYBBP1A was synthesized using in vitro transcription/translation-coupled reticulocyte lysate system (Promega). Binding was performed in TNE buffer [150 mM NaCl, 0.5% Nonidet P-40, 50 mM Tris-HCl (pH 8.0), 5 mM EDTA] for 30 min under rotation at 4°C, and the beads were washed 5 times with TNE buffer. Beads were boiled in SDS sample buffer for 5 min, and the supernatants were loaded to SDS-polyacrylamide gels followed by autoradiography.

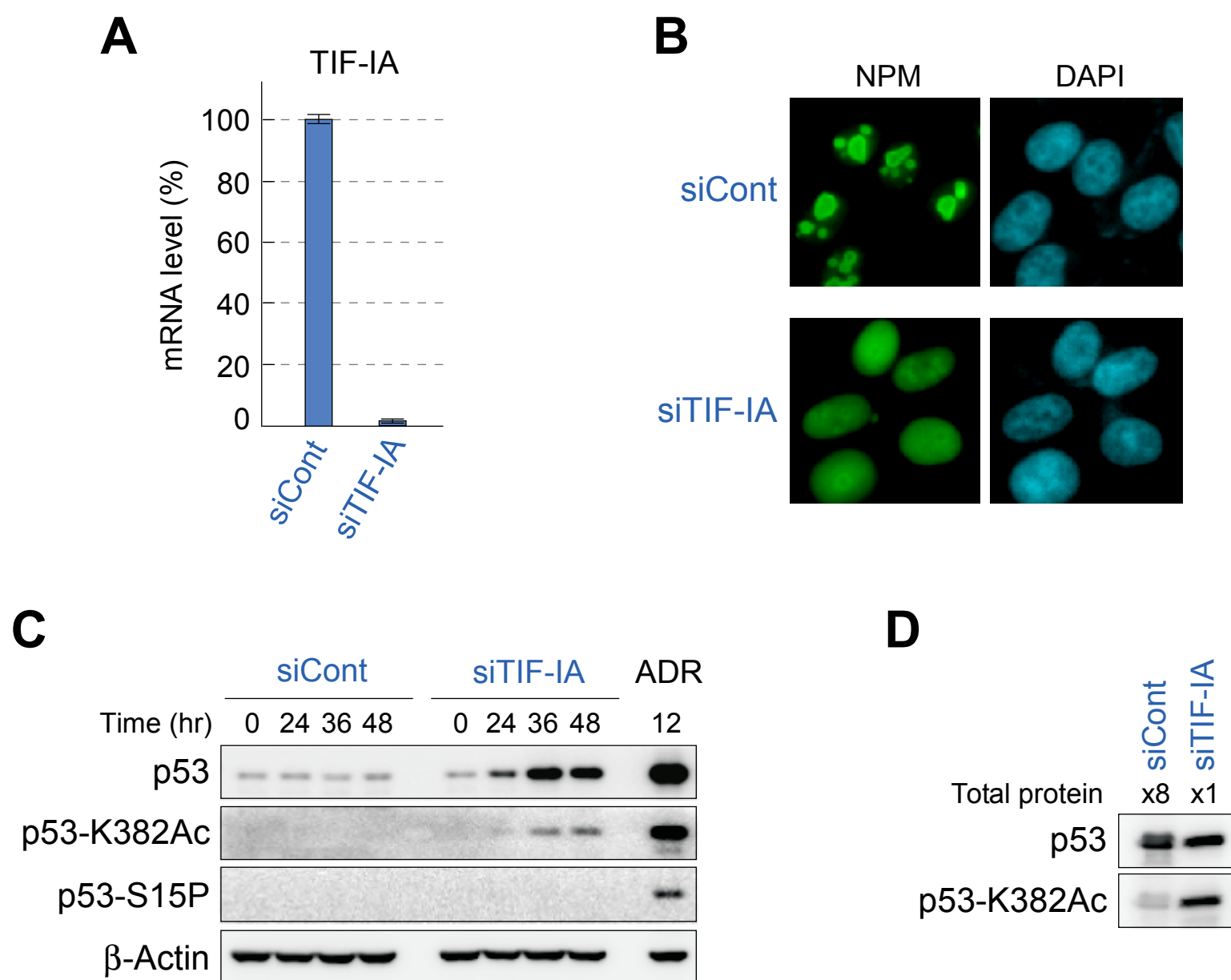
Nuclear run-on assay

cDNAs probes corresponding to 28S rRNA was amplified by polymerase chain reaction, cloned into pGEM-T easy vector and spotted onto Hybond-N+ membrane (GE Healthcare). Run-on assay was performed on MCF-7 cells transfected with Control, MYBBP1A, RPL5, or RPL11 siRNA with or without TIF-IA siRNA. Approximately 1.0×10^7 cells were collected on ice and suspended in lysis buffer [10 mM Tris-HCl (pH 7.4), 10 mM NaCl, 3 mM MgCl₂ and 0.5% NP-40] for 10 min. Nuclei were collected

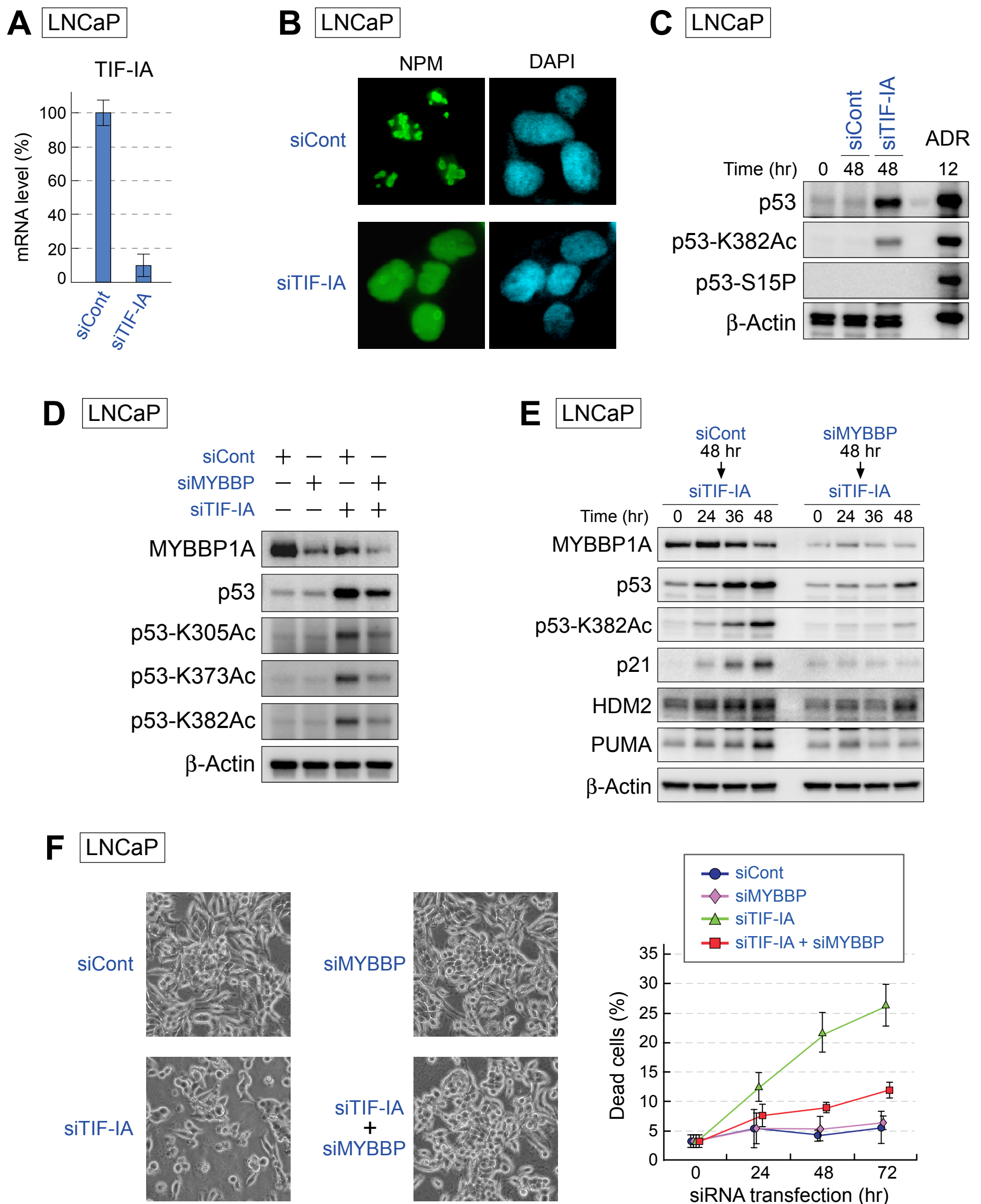
by centrifugation at $1,000 \times g$ for 5 min, and washed twice with the same buffer. The nuclei were suspended in 50 mM Tris-HCl (pH 8.3), 40% glycerol, 5 mM $MgCl_2$ and 0.1 mM EDTA. The nuclei were mixed with an equal volume of reaction buffer [50 mM Tris-HCl (pH 8.0), 5 mM $MgCl_2$, 300 mM KCl, 0.5 mM of ATP, CTP, GTP and 100 μ Ci of α - ^{32}P UTP] and incubated for 30 min at 30°C. Nuclear RNA was extracted and resuspended in RapidHyb hybridization buffer (GE Healthcare). Hybridization was carried out at 42°C for 24 hour. Membrane was washed twice at room temperature in 2 \times SSC containing 0.1% SDS and twice at 65°C in 0.2 \times SSC containing 0.1% SDS, and then exposed on BioMAX MS film.



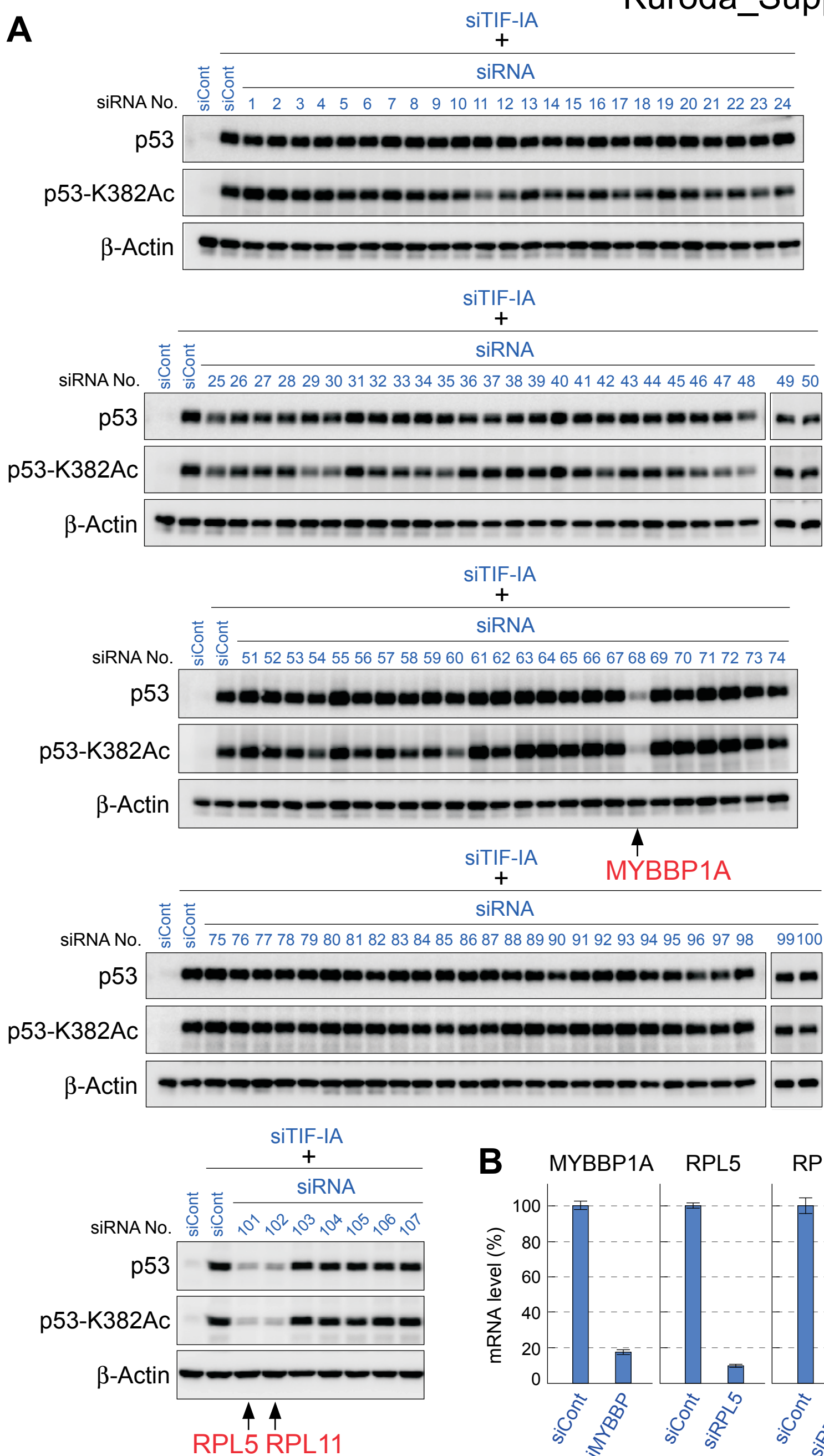
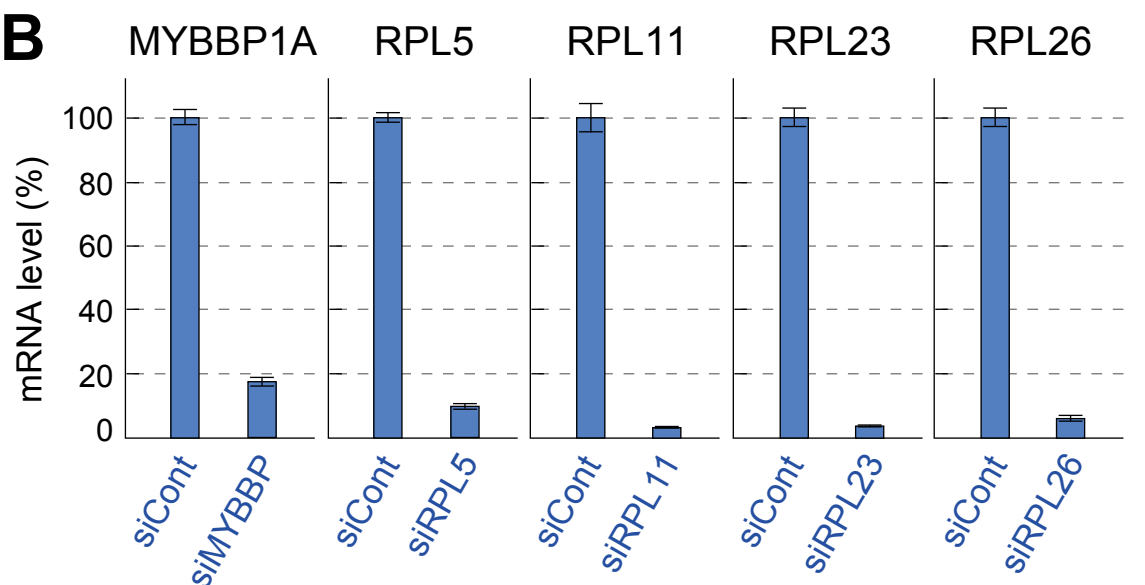
Supplementary Figure S1 Nucleolar disruption induced by low dose of ActD treatment causes acetylation of p53 protein without phosphorylation. **(A)** Low dose of ActD treatment induced accumulation and acetylation but not phosphorylation of p53 protein. MCF-7 cells were treated with a low concentration (5 nM) of ActD for indicated times, and the cell lysates were analyzed by immunoblot using the indicated antibodies. Lysates prepared from ADR (0.5 μ g/ml)-treated cells were used as a positive control. **(B)** Nucleolar structure was disrupted by low dose of ActD treatment. MCF-7 cells were treated with 5 nM ActD for 8 hours, and the localization of NPM was visualized by immunofluorescence.



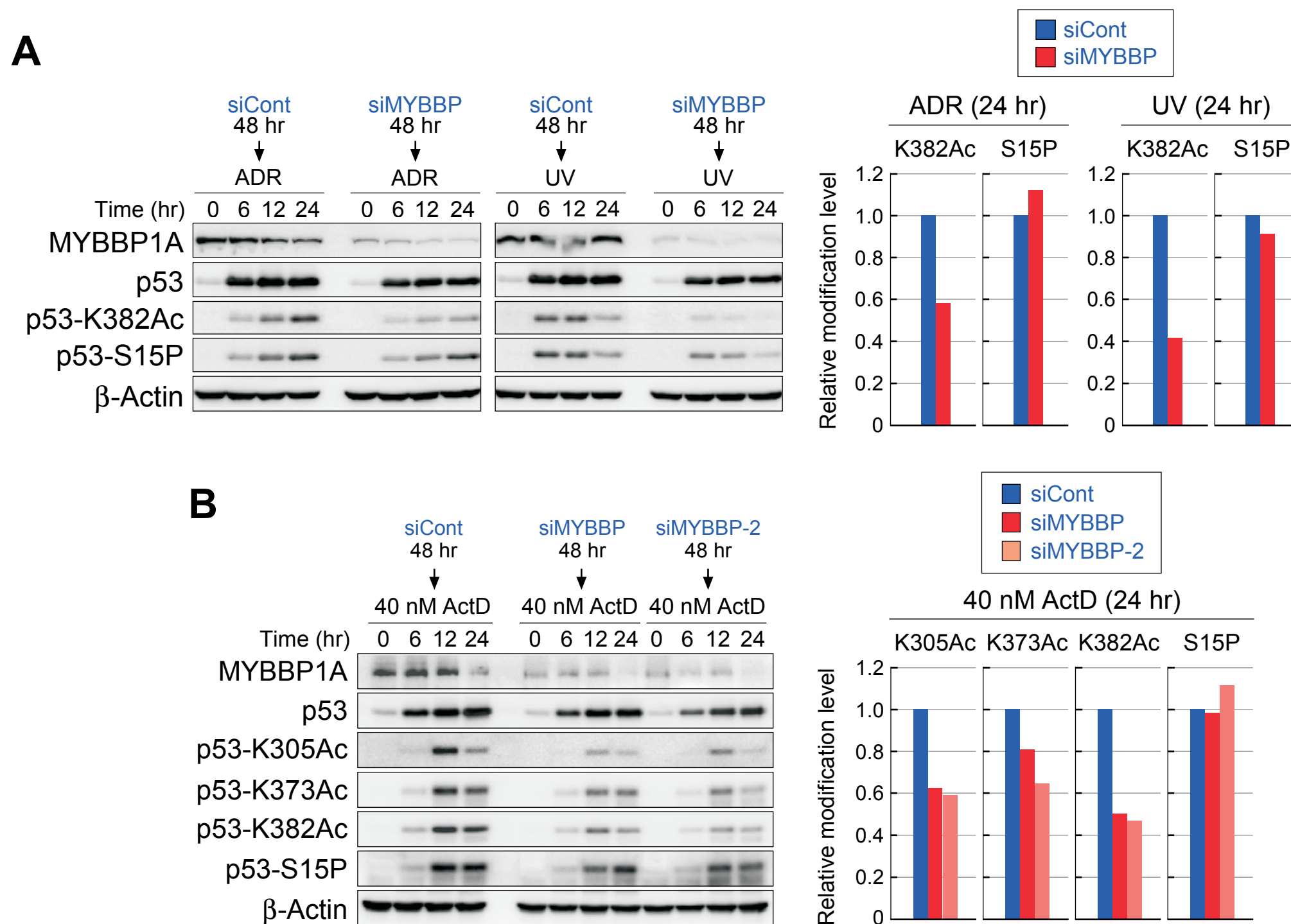
Supplementary Figure S2 Nucleolar disruption induced by TIF-IA knockdown causes acetylation of p53 protein without phosphorylation. **(A)** Knockdown efficiency of TIF-IA siRNA. MCF-7 cells were treated with siCont or siTIF-IA for 48 hours, and the mRNA levels were determined by RT-qPCR. The mRNA levels were normalized to β -Actin. Values are given as the mean \pm SD for triplicate experiments. **(B)** Nucleolar structure was disrupted in TIF-IA siRNA treated cells. MCF-7 cells were treated with siCont or siTIF-IA for 48 hours, and the localization of NPM was visualized by immunofluorescence using an anti-NPM antibody. **(C)** Knockdown of TIF-IA induced accumulation and acetylation but not phosphorylation of p53 protein. MCF-7 cells were treated with siCont or siTIF-IA for the indicated times, and the cell lysates were analyzed by immunoblot using the indicated antibodies. Lysates prepared from ADR (0.5 μ g/ml)-treated cells were used as a positive control. **(D)** Knockdown of TIF-IA induced acetylation of p53 protein. MCF-7 cells were treated with siCont or siTIF-IA for 48 hours. An equal amount of p53 was loaded, and acetylation levels were analyzed by immunoblot using an anti-acetylated p53 antibody.



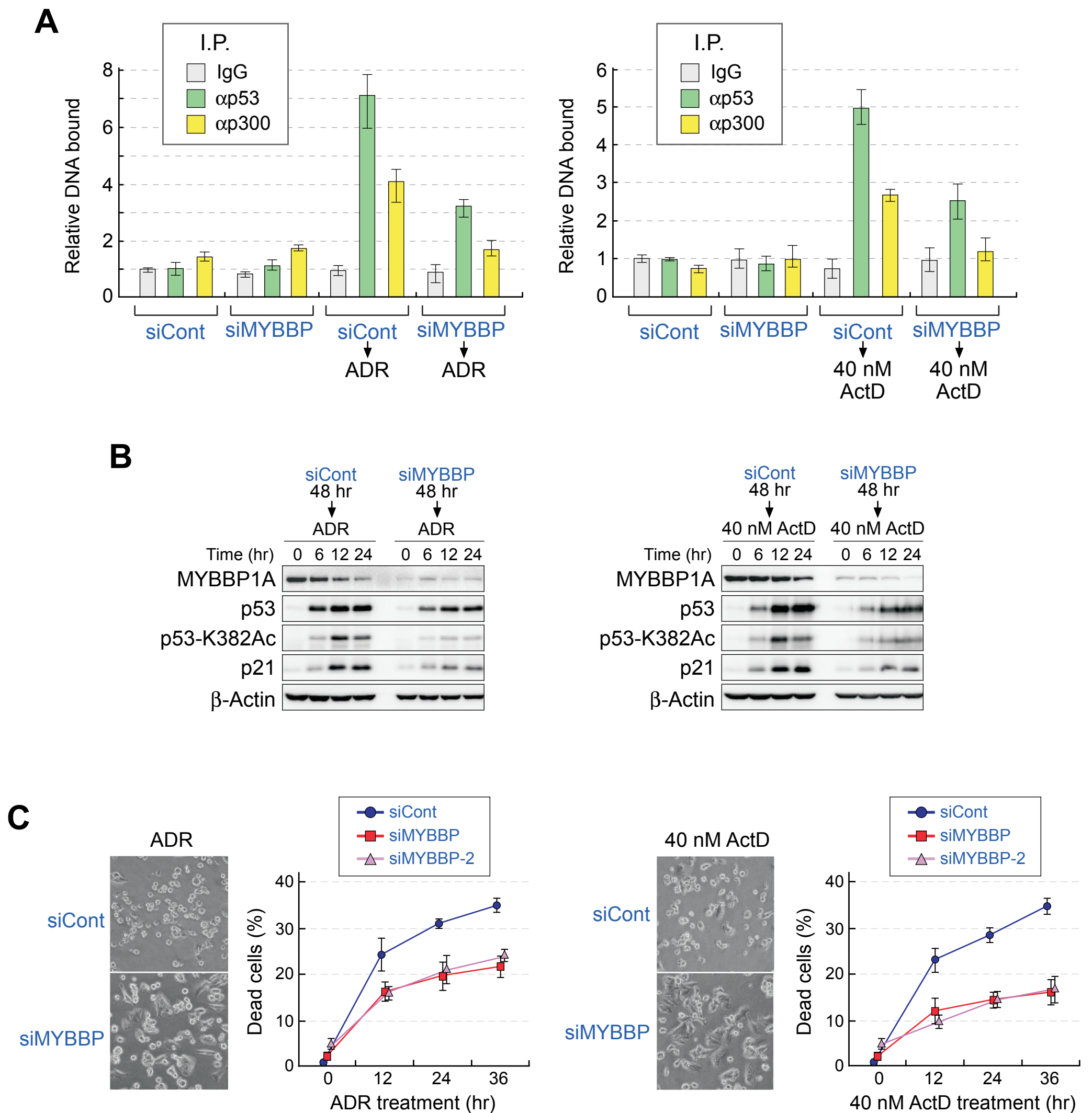
Supplementary Figure S3 MYBBP1A is necessary for the acetylation and transactivation of p53 and nucleolar stress-induced apoptosis in LNCaP cells. **(A)** Knockdown efficiency of TIF-IA siRNA in LNCaP cells. LNCaP cells were treated with siCont or siTIF-IA for 48 hours, and the mRNA levels were determined by RT-qPCR. **(B)** Immunofluorescence analysis showed translocation of NPM in LNCaP cells treated with siTIF-IA for 48 hours. **(C)** Knockdown of TIF-IA induced accumulation and acetylation but not phosphorylation of p53 protein in LNCaP cells. Lysates prepared from ADR (0.5 μ g/ml)-treated cells were used as a positive control. **(D)** Knockdown of MYBBP1A decreased the acetylation levels at multiple lysine residues in p53 protein in LNCaP cells treated with siTIF-IA. **(E)** Knockdown of MYBBP1A reduced the elevation of p53 target gene products induced by siTIF-IA in LNCaP cells. **(F)** Knockdown of MYBBP1A decreased the level of apoptosis induced by siTIF-IA in LNCaP cells. Left: the phase-contrast images of LNCaP cells treated with the indicated siRNAs for 72 hours. Right: the percentage of dead cells was measured by trypan blue exclusion assay. Values are given as the mean \pm SD for triplicate experiments.

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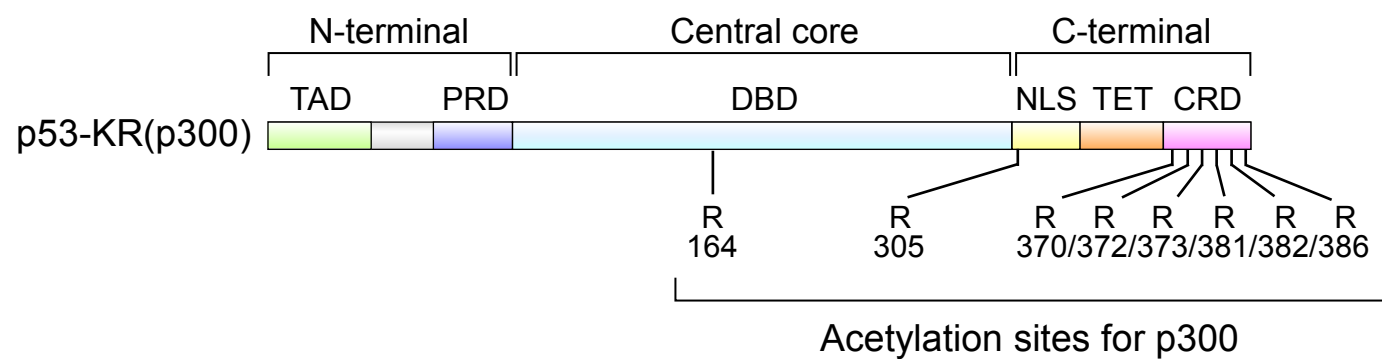
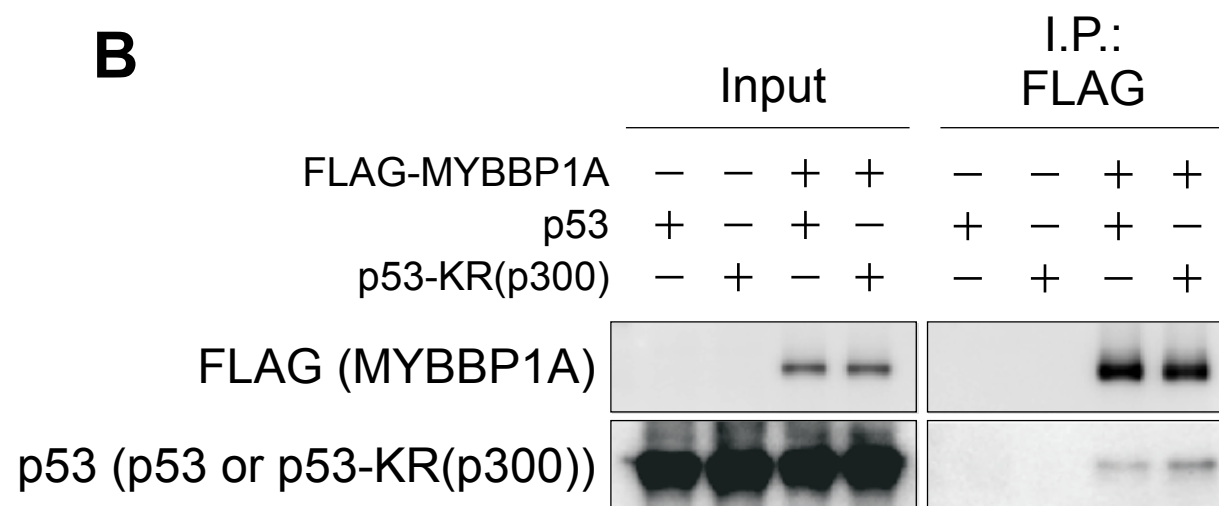
Supplementary Figure S4 Identification of nucleolar proteins that are involved in p53 acetylation and accumulation. **(A)** MCF-7 cells were treated with each siRNA and siTIF-IA for 48 hours, and the cell lysates were analyzed by immunoblot using anti-p53 and anti-acetylated p53 antibodies. #68 is siRNA against MYBBP1A. #101 and #102 are siRNAs against RPL5 and RPL11, respectively. **(B)** Knockdown efficiency of MYBBP1A, RPL5, RPL11, RPL23, or RPL26 siRNA, respectively. MCF-7 cells were treated with the indicated siRNAs for 48 hours, and the mRNA levels were determined by RT-qPCR. The mRNA levels were normalized to β-Actin. Values are given as the mean \pm SD for triplicate experiments.



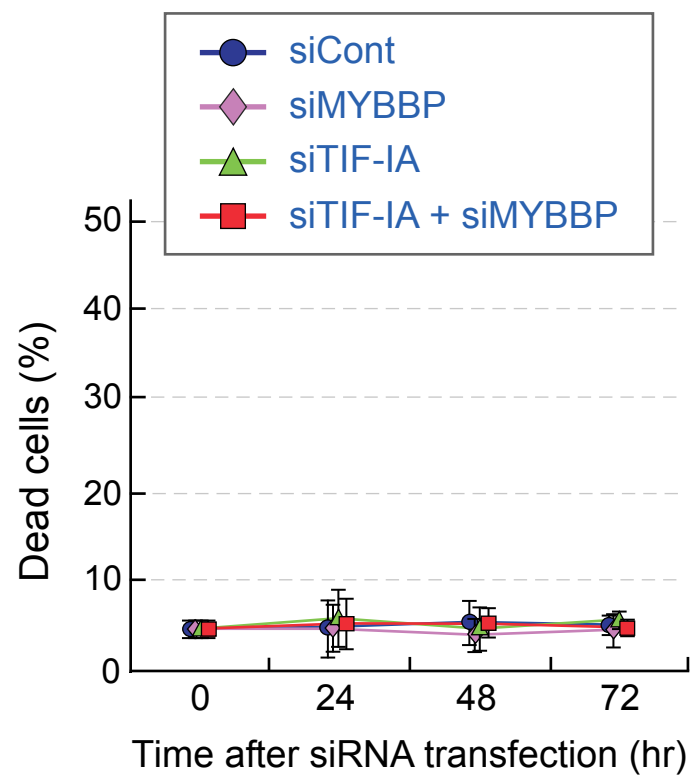
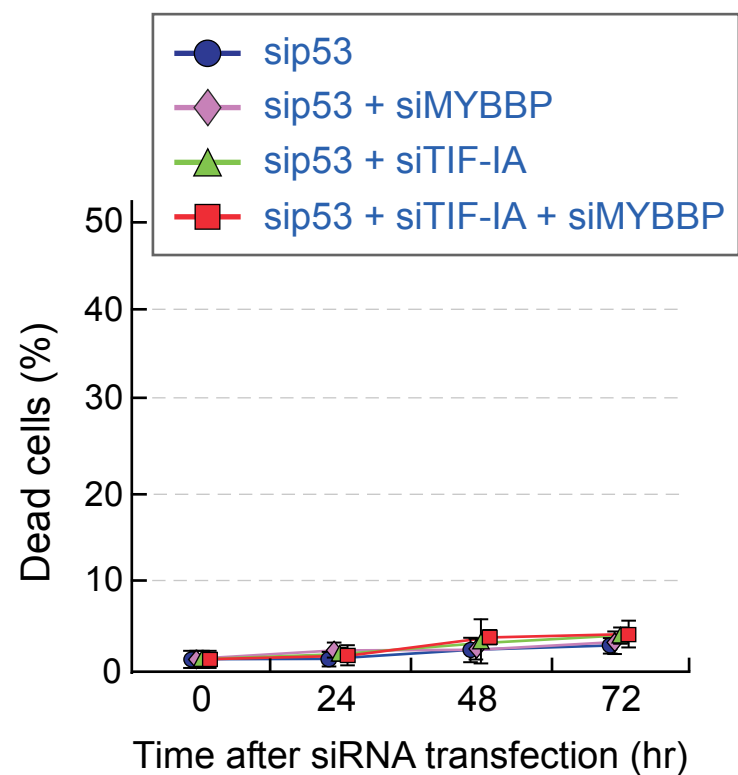
Supplementary Figure S5 MYBBP1A is necessary for the acetylation and accumulation of p53 induced by nucleolar stress. **(A)** Knockdown of MYBBP1A decreased p53 acetylation levels induced by ADR or UV treatment. Left: MCF-7 cells were treated with the siCont or siMYBBBP for 48 hours. The cells were then treated with ADR (0.5 $\mu\text{g}/\text{ml}$) or UV (25 J/m^2) for the indicated times. The cell lysates were analyzed by immunoblot using the indicated antibodies. Right: relative quantification of acetylation and phosphorylation levels of p53 protein. Twenty-four hours after ADR or UV treatment, the intensity of the acetylated or phosphorylated p53 proteins were quantified by phosphoimager analysis and plotted. The intensities of the modified p53 proteins were corrected using p53 protein level. The intensity of the siCont-treated cells was normalized to 1.0. **(B)** Knockdown of MYBBP1A decreased the acetylation levels at multiple lysine residues in p53 protein by 40 nM ActD treatment. Left: MCF-7 cells were treated with siCont or two independent siMYBBBP (siMYBBBP and siMYBBBP-2) for 48 hours, followed by 40 nM ActD treatment at the indicated times. The cell lysates were analyzed by immunoblot using the indicated antibodies. Right: relative quantification of acetylation and phosphorylation levels of p53 protein. Twenty-four hours after 40 nM ActD treatment, the intensity of the modified p53 proteins were quantified as (A).



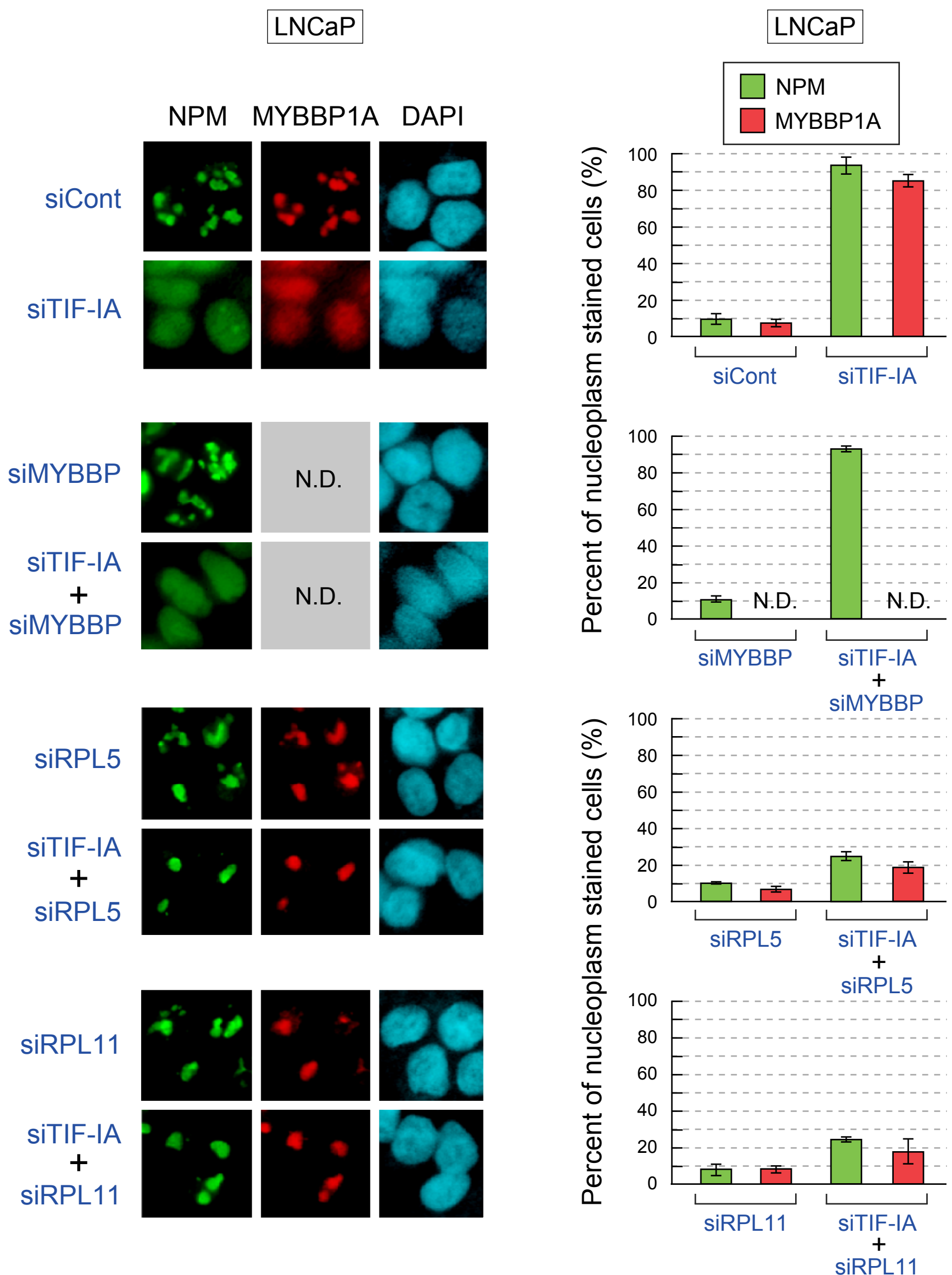
Supplementary Figure S6 MYBBP1A is necessary for the acetylation and transactivation of p53 and apoptosis induced by ADR or ActD. **(A)** Knockdown of MYBBP1A reduced the recruitment of p53 and p300 to the *p21* promoter in the ADR- or ActD-treated cells. MCF-7 cells were treated with siCont or siMYBBP for 48 hours prior to treatment with ADR (0.5 $\mu\text{g/ml}$) for 12 hours or ActD (40 nM) for 24 hours. A ChIP assay was performed using normal rabbit IgG, anti-p53, or anti-p300 antibodies. The p53-binding region of the *p21* promoter was amplified and analyzed by qPCR. Values are given as the means \pm SD for triplicate experiments. **(B)** Knockdown of MYBBP1A reduced the elevation of p21 protein levels induced by ADR or ActD treatment. MCF-7 cells were treated with siCont or siMYBBP for 48 hours prior to treatment with ADR (0.5 $\mu\text{g/ml}$) or ActD (40 nM) for the indicated times. The cell lysates were prepared and analyzed by immunoblot using the indicated antibodies. **(C)** Knockdown of MYBBP1A decreased the level of apoptosis induced by ADR or ActD treatment. Left photos: the phase-contrast images of MCF-7 cells treated with the indicated siRNAs for 48 hours, and then treated with ADR (0.5 $\mu\text{g/ml}$) or ActD (40 nM) for 36 hours. Representative images are shown. Right graphs: percentage of dead cells. The percentage of dead cells was measured by trypan blue exclusion assay. Values are given as the mean \pm SD for triplicate experiments.

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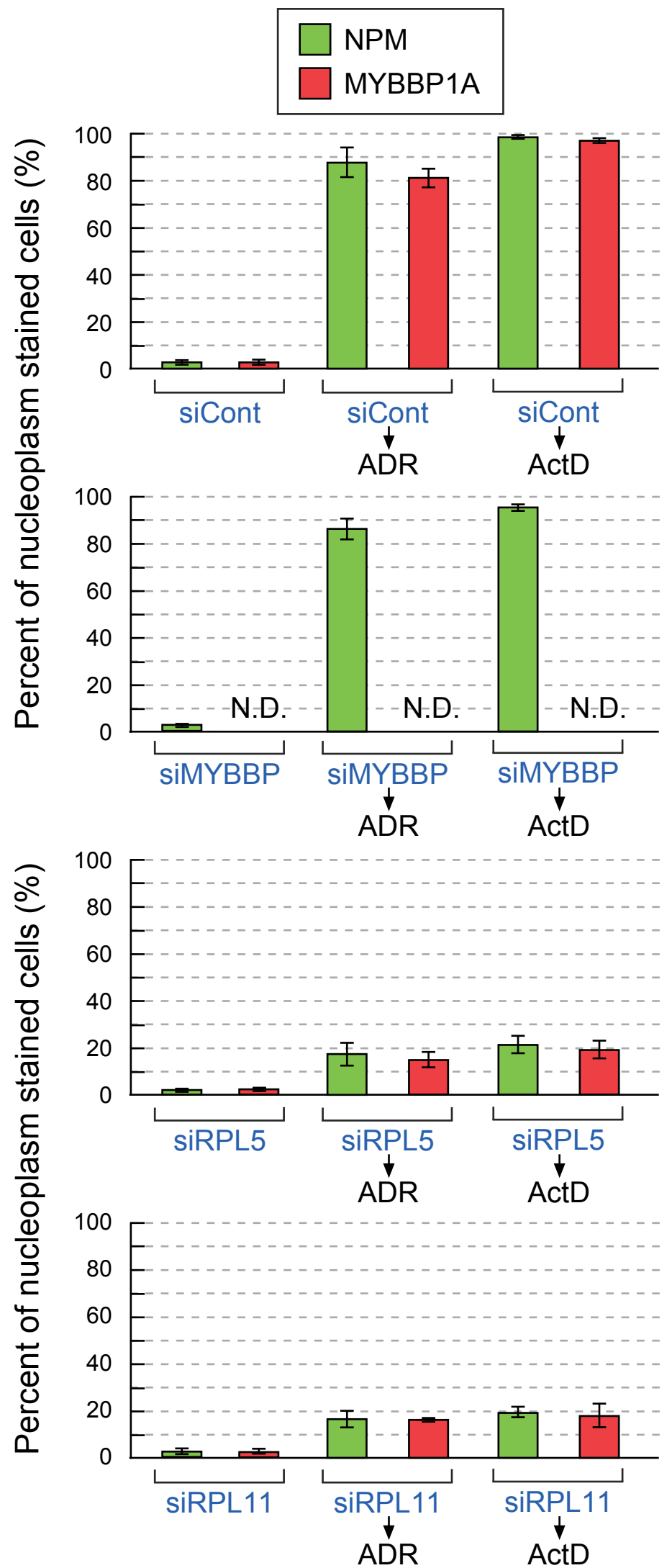
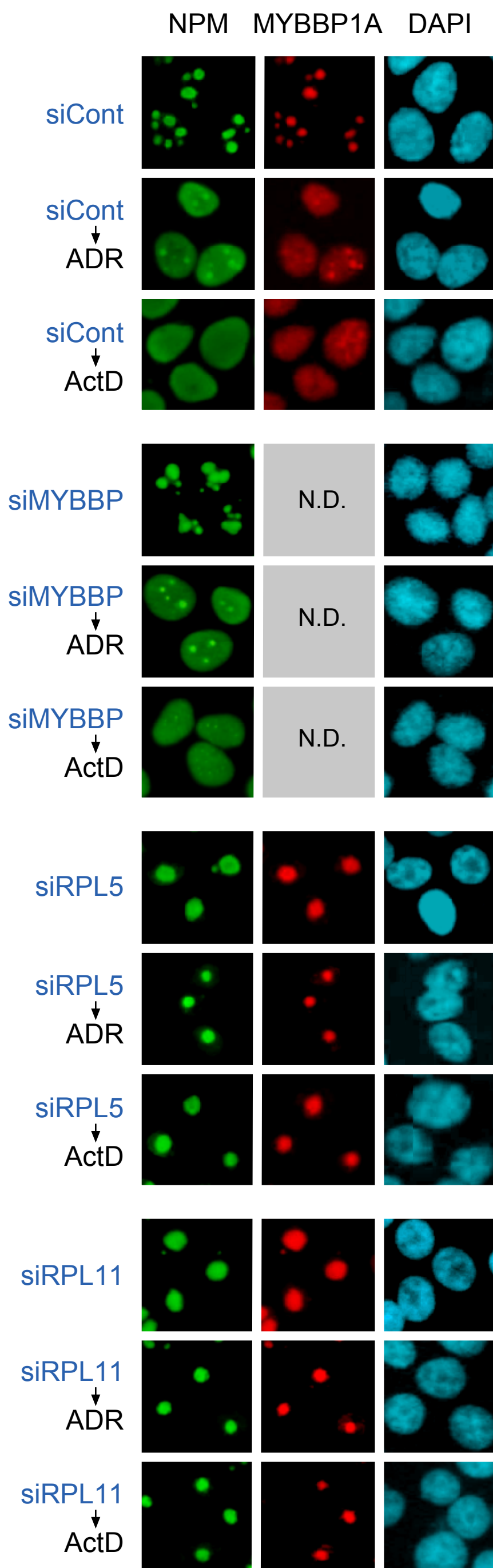
Supplementary Figure S7 MYBBP1A interacts with p53-KR(p300). **(A)** Schematic representation of the p53-KR(300) mutant. We generated p53 lysine-to-arginine mutant at eight acetylation sites for p300. **(B)** MYBBP1A interacted with p53-KR(p300). H1299 cells were transfected with combination of the expression vectors for FLAG-MYBBP1A, p53, and/or p53-KR(300), as indicated. Twenty-four hours after transfection, the cells were treated with ADR (0.5 μ g/ml) for 12 hours. MYBBP1A was immunoprecipitated from the cell lysates using an anti-FLAG antibody and p53 association was detected by immunoblot using an anti-p53 antibody.

A H1299**B** MCF-7

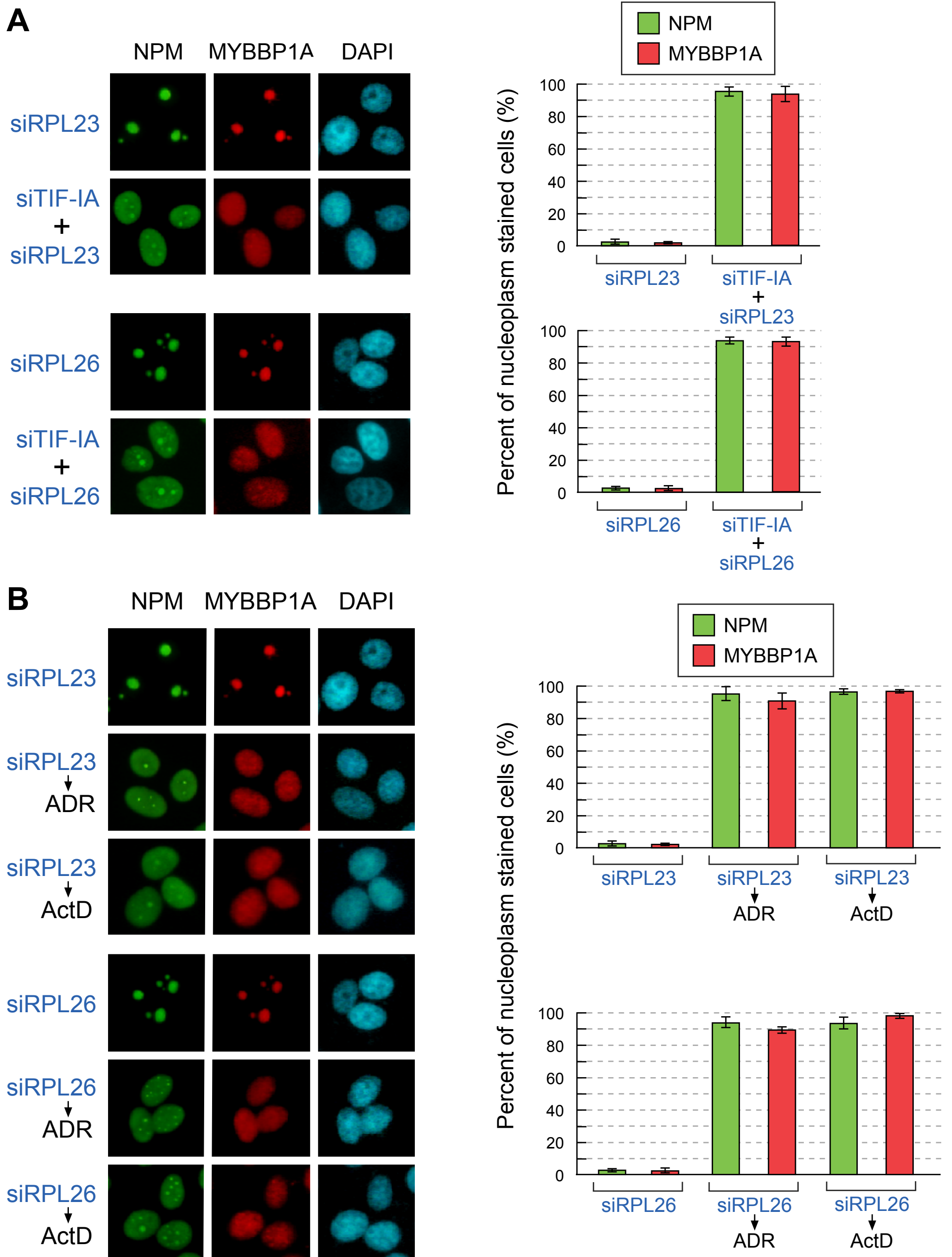
Supplementary Figure S8 Apoptosis induced by TIF-IA knockdown is dependent on the presence of p53. (A) The percentage of dead H1299 cells. H1299 cells were transfected with the indicated siRNAs for the indicated times, and the percentage of dead cells was measured by trypan blue exclusion assay. Values are given as the mean \pm SD for triplicate experiments. (B) The percentage of dead p53-depleted MCF-7 cells. MCF-7 cells were treated with sip53 for 48 hours prior to treatment with the indicated siRNAs for the indicated times. The percentage of dead cells was measured as (A).



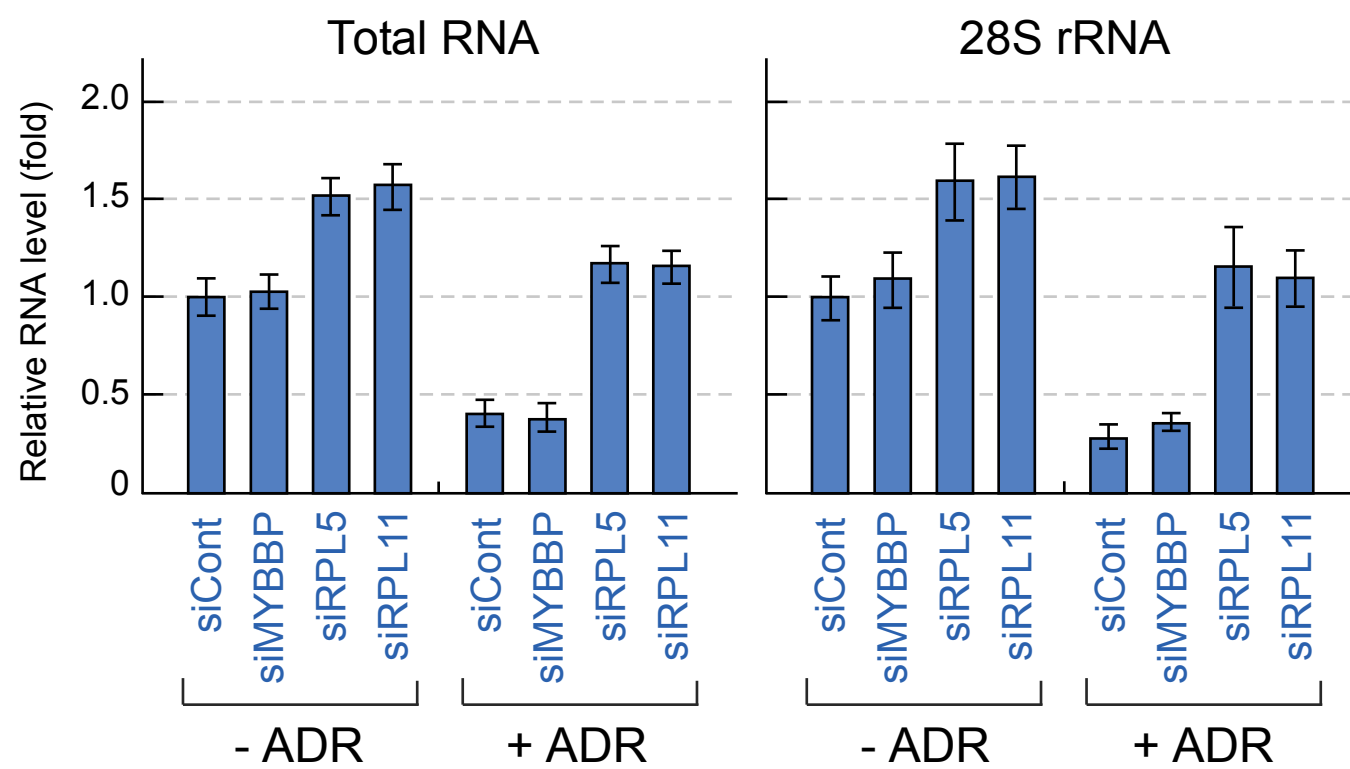
Supplementary Figure S9 RPL5 and RPL11 are required for the MYBBP1A translocation from the nucleolus in LNCaP cells. MYBBP1A translocation from the nucleolus to the nucleoplasm induced by TIF-IA knockdown was hampered by the knockdown of RPL5 or RPL11. Left: LNCaP cells were treated with the indicated siRNAs for 48 hours, and the cells were stained with the indicated antibodies or DAPI. N.D. means “not determined”. Right: The percentage of the cells that showed translocation of NPM or MYBBP1A from the nucleolus after treatment with the indicated siRNAs. Values are given as the mean \pm SD for triplicate experiments.



Supplementary Figure S10 RPL5 and RPL11 are required for the MYBBP1A translocation from the nucleolus in ADR or ActD treated cells. MYBBP1A translocation from the nucleolus to the nucleoplasm induced by ADR or ActD treatment was hindered by the knockdown of RPL5 or RPL11. Left: MCF-7 cells were treated with the indicated siRNAs for 48 hours, followed by the treatment with ADR (0.5 μ g/ml) or ActD (40 nM) for 24 hours. The cells were stained with the indicated antibodies or DAPI. N.D. means "not determined". Right: The percentage of the cells that showed translocation of NPM or MYBBP1A from the nucleolus after the treatment. Values are given as the mean \pm SD for triplicate experiments.



Supplementary Figure S11 RPL23 and RPL26 are not required for the MYBBP1A translocation induced by nucleolar disruption. **(A)** Left: MCF-7 cells were treated with the indicated siRNAs for 48 hours, and the cells were stained with the indicated antibodies or DAPI. Right: The percentage of cells that showed translocation of NPM or MYBBP1A from the nucleolus after treatment with the indicated siRNAs. Values are given as the means \pm SD for triplicate experiments. N.D. means "not determined". **(B)** Left: MCF-7 cells were treated with the indicated siRNAs for 48 hours, followed by the treatment with ADR (0.5 μ g/ml) or ActD (40 nM) for 24 hours. The cells were stained with the indicated antibodies or DAPI. Right: The percentage of cells that showed translocation of NPM or MYBBP1A from the nucleolus after treatment with the indicated siRNAs. Values are given as the mean \pm SD for triplicate experiments. N.D. means "not determined".



Supplementary Figure S12 Knockdown of RPL5 or RPL11 retains RNA content in the nucleolus in ADR treated cells. Total RNA was isolated from the isolated nucleoli of MCF-7 cells transfected with the indicated siRNAs for 48 hours and then treated with ADR (0.5 μ g/ml) for 24 hours. The total RNA (Left) and 28S rRNA (Right) levels were quantified by spectrophotometry and RT-qPCR, respectively.

Kuroda_Supplementary Table S1

Supplementary Table S1 List of the nucleolar proteins that were knocked-down in the screen

siRNA No.	Name	Sequence	
1	DNTTIP2	GGAAGAACUGCUGGCUGAUUCUGAA UUCAGAAUCAGCCAGCAGUUCUUC	sense antisense
2	WDSOF1	AUAAGAGUCCUCUCGUUGGUCCCGG CCGGGACCAACGAGAGGACUCUUAU	sense antisense
3	RPS19BP1	UACUCUGAGGUGGUCUCGACACUCU AGAGUGUCGAGACCACCUCAGAGUA	sense antisense
4	RFC2	UCCUCACCAGGCUGAUGAAUGUUAU AUAACAUAUCAGCCUGGUGAGGA	sense antisense
5	DIMT1L	CCCGUCCAUGGACAUAGAUGACUU AAGUCAUCUAUGUCCAUGGAACGGG	sense antisense
6	EXOSC3	UCACUAUGCCAAUCACAUGGUCUCC GGAGACCAUGUGAUUGGCAUAGUGA	sense antisense
7	CDKN2AIP	AAUAGUGGCAAUUCUACCUCUGAGC GCUCAGAGGUAGAAUUGCCACUAUU	sense antisense
8	RFC1	CCCAUCCCUGGAUUCGGAAUACAAU AUUGUAUUCGAAUCCAGGGAUGGG	sense antisense
9	TSPYL1	GGUGUCUCUUUCUACUCCAAUUAUA UAUAAUUGGAGUAGAAAGAGACACC	sense antisense
10	PRKDC	UAAACACGGUCGUCACCAGUGUCUG CAGACACUGGUGACGACCGUGUUUA	sense antisense
11	DDX56	UAACCGGGUUAUGUAAUAUCAGCUC GAGCUGAUUAUCAUAACCCGGUUA	sense antisense
12	EXOSC4	CCCACGCUCCAGAUUGAUUAUCUAU AUAGAUUAUCAUUCUGGGAGCGUGGG	sense antisense
13	DIS3	CAGCAGACAACCAGCUGCAAGUUUAU AUAACUUGCAGCUGGUUGUCUGCUG	sense antisense
14	C6orf153	UUGGCUCUGCAAUCCAACAGCUCC GGAGCUGUUGGAAUUGCAGAGCCAA	sense antisense
15	SSB	AUACUCAUCAGUCACUUCAGGUAGG CCUACCUGAAGUGACUGAUGAGUAU	sense antisense
16	CCDC59	GAGCCUUUAUUUGAAGAUCAGUGUA UACACUGAUCUCAAUAAGGCUC	sense antisense

17	REXO4	CAUAGGUCCAGAGGGCGCCAAGAU UAUCUUGGCCGCCUCUGGACCUAUG	sense antisense
18	PINX1	CCACAGGUAAGAUGUGGAAAGUUA UAACUUUCCACAUCUUUACCUGUGG	sense antisense
19	KIF2C	GGGCAGACAUUUGCCAACUCCA AAUUGGAGUUGGCAAUGUCUGCCC	sense antisense
20	PTBP1	AGAAGGACCGCAAGAUGGCACUGAU AUCAGUGCCAUCUUGCGGUCCUUCU	sense antisense
21	PARP1	AGAAGCUGCAGCUUGUAGUAGGAGU ACUCCUACUACAAGCUGCAGCUUCU	sense antisense
22	XRCC1	UUAUCUCGCAGCUCGGAGCGGAAGG CCUCCGCUCCGAGCUGCGAGAUAA	sense antisense
23	EMG1	UUGAGUAGCUCAUAUGUCUUCCCUA UAGGGAAGACAUUAGAGCUACUCAA	sense antisense
24	ABT1	UUUACGGGCCUCAGUUCUGUCUCA UGAGCAGGAACUGAGGGCCCGUAAA	sense antisense
25	SMC4	CAAGGACAUUCAGAGUUGUACAGUA UACUGUACAACUCUGAAUGUCCUUG	sense antisense
26	SKIV2L2	GGGAAUUAACAUGCCAGCUAGAACU AGUUCUAGCUGGCAUGUAAUUC	sense antisense
27	PNO1	AAGAUUAGGUUGCAAUUGGCAGUUC GAACUGCCAUUUGCAACCUAAUCUU	sense antisense
28	C14orf21	UGUUCGUGGACAAAGCUAGGGCCUG CAGGCCCUAGCUUUGUCCACGAACA	sense antisense
29	NCL	UUCUGUUGCACUGUAGGAGAGGUUG CAACCUCUCCUACAGUGCAACAGAA	sense antisense
30	PRPF19	UUUCAGGGUAGCCAGAGCUUCUCGG CCGAGAAGCUCUGGCUACCCUGAAA	sense antisense
31	XRCC5	UUGGAAACAAGUCUUAAGGGUGUC GACACCCUUGAAGACUUGUUUCCAA	sense antisense
32	CDC2L5	GCCUCCCAUGCUGCCUGAAGAUAAA UUUAUCUUCAGGCAGCAUGGGAGGC	sense antisense
33	FCF1	CAAGGAUCCAAGAUUUGAACGAUUA UAAUCGUUCAAAUCUUGGAUCCUUG	sense antisense
34	BYSL	CAUUGUGGCAGCCUUCUCCAGGGUG CACCCUGGAGAAGGCUGCCACAAUG	sense antisense

35	BLM	AACAGAAGGAAACUUCUGGCGAAGC GCUUCGCCAGAAGUUUCCUUCUGUU	sense antisense
36	UTP18	GCUCAUAAAUGGCAUUGCUGGAUUAU AUAUCCAGCAAUGCCAUUUAUGAGC	sense antisense
37	ADAR	CAGAGAGGUGUUGAUUGCCUUUCCU AGGAAAGGCAAUCAACACCUCUCUG	sense antisense
38	POP1	UGAAGUUC CAGGAUGAGGCUCUUGU ACAAGAGCCUCAUCCUGGAACUUCA	sense antisense
39	RCL1	CAACAAGUUCAUACCUGAUUAUCUAU AUAGAUUAUCAGGUAUGAACUUGUUG	sense antisense
40	CD3EAP	UAGAGGCUCAGUGUUAUUCUGUUC GGAACAGAUUAACACUGAGCCUCUA	sense antisense
41	NCAPD3	CGGAGUCAAGGAAAUGACAUCUUAU AUAAGAUGUCAUUUCCUUGACUCCG	sense antisense
42	WDR3	GGGACAUAGCUUAUCUGCAAGAGAU AUCUCUUGCAGAUAAAGCUAUGUCCC	sense antisense
43	RFC5	CCAACAUCCUGGACUGGAUGUUGAA UUCAACAUCCAGUCCAGGAUGUUGG	sense antisense
44	RBM39	CAAGAGCCCUGUGAGAGAACCUAUU AAUAGGUUCUCUCACAGGGCUCUUG	sense antisense
45	AURKB	CCCGACAUCUUAACGCGGCACUUCA UGAAGUGCCGCGUUAAGAUGUCGGG	sense antisense
46	PAI-RBP1	UAAACACACUGAUUUAGGAGUGCGU ACGCACUCCUAAAUCAGUGUGUUUA	sense antisense
47	SMC2	AUUAGAAGCCCGAACCCUGAGACAGG CCUGUCUCAGGUUCGGGCUUCUAAU	sense antisense
48	MKI67IP	UUUAAAUCGCUCCUCCAUCCGUAGC GCUACGGAUGGAGGAGCGAUUUAAA	sense antisense
49	ISG20L2	UUUAUAGCUCCAUGGUGGCCUGGG CCCAGGCCACCAUGGAGCUAUUAAA	sense antisense
50	TBL3	UCAGCUGAUGGCACCAUCAAGCUCU AGAGCUUGAUGGUGCCAUCAGCUGA	sense antisense
51	ILF3	CCACUGAUGCUAUUGGGCAUCUAGA UCUAGAUGCCCAAUAGCAUCAGUGG	sense antisense
52	ESF1	UUUGCAACCCGUCUAAACCGCUGGU ACCAGCGGUUUAGACGGGUUGCAAA	sense antisense

53	TOP1	GGGAAGGACUCCAUCAGAUACUAUA UAUAGUAUCUGAUGGAGUCCUUC	sense antisense
54	BMS1	CAGUGCAGAGGAAGAAGACUCAGAA UUCUGAGUCUUCUCCUCUGCACUG	sense antisense
55	SRP14	UUCAAGGUGAUUAUAGACGCUGCCCG CGGGCAGCGUCUAUAUCACCUUGAA	sense antisense
56	FBXL11	GAACCCGAAGAAGAAAGGAUUCGUU AACGAAUCCUUCUUCUUCGGGUUC	sense antisense
57	GPATCH4	AUAACGGGUGGUCUCCUAGAAAGG CCUUUCUAAGGAGACCACCCGUUAU	sense antisense
58	HIST1H1C	UUUCGUUUGCACCAGAGUGCCCUUG CAAGGGCACUCUGGUGCAAACGAAA	sense antisense
59	SEN3	CCCUCAGCACUGAUGAGGUAGUAGA UCUACUACCUCUACAGUGCUGAGGG	sense antisense
60	H2AFZ	UGGGCCGUUUCAUCGACACCUAAA UUUAGGUGUCGAUGAAUACGGCCCA	sense antisense
61	RSL1D1	UUAUCCAGCUGCUUCCGUGCUGUCG CGACAGCACGGAAGCAGCUGGAUAA	sense antisense
62	ZCCHC9	UUAUAUGUGGUACUAACUCGGGCC GGGCCCGAGUUAGUACCACAUUAA	sense antisense
63	VRK1	AAGAGGUCCAUUGUCACUGGGUUC GGAACCCAGUGACAAUGGACCUCUU	sense antisense
64	CDK7	CCAAACUGUCCAGUGGAAACCUUAA UUAAGGUUCCACUGGACAGUUUGG	sense antisense
65	WDR36	CCGGACUGACAUUUCUCCAUAAGAGA UCUCUAUGGAGAAAUGUCAGUCCGG	sense antisense
66	CDK9	CAGAUGCUGCUUAACGGCCUCUACU AGUAGAGGCCGUUAAGCAGCAUCUG	sense antisense
67	NOC4L	UGUCGUCGUAGUCCAGGUACUCCCG CGGGAGUACCUGGACUACGACGACA	sense antisense
68	MYBBP1A	UUCACCAGCCGACCUGACUGAAAGA UCUUUCAGUCAGGUCGGCUGGUGAA	sense antisense
69	WDR18	UGGACCUGGCUGAGCACCAUAUGUU AACAUUAGGUGCUCAGCCAGGUCCA	sense antisense
70	WDR75	UUGCAAGCCAGUUACCAAGCAGCC GGCUGCUUUGGUAACUGGCUUGCAA	sense antisense

71	TOP2A	GCUCAGCUCUUUGGCUCGAUUGUUA UAACAAUCGAGCCAAAGAGCUGAGC	sense antisense
72	CIRH1A	CAGGAAUCCGCUGUGUGGCUUACAA UUGUAAGCCACACAGCGGAUCCUG	sense antisense
73	NOL14	AUCAAUGACUGUGAGGUCUGGUGGG CCCACCAGACCUCACAGUCAUUGAU	sense antisense
74	C1orf107	UUAGGGUGUUGAGUAGCUGGCUCUG CAGAGCCAGCUACUCAACACCCUAA	sense antisense
75	TWISTNB	CCUUAACAGGAAACGCACCGGCAUU AAUGCCGGUGCGUUUCCUGUUAAGG	sense antisense
76	KRR1	UUCUGAUUCAUCUUGGUUCUCCGGC GCCGGAGAACCAAGAUGAAUCAGAA	sense antisense
77	TEX10	UUCAGUAGAUGACAGGCCUUCAUCC GGAUGAAGGCCUGUCAUCUACUGAA	sense antisense
78	TTF1	GGAUGACUAAUGGUCGGCGUAUCUA UAGAUACGCCGACCAUAGUCAUCC	sense antisense
79	BOP1	UGUAGAUGC GCCUGCCAUCCAGGUC GACCUGGAUGGCAGGCGCAUCUACA	sense antisense
80	POLR1A	CCAUCUCUCCCUGGUUGCUGAUUUAU AUAAUCAGCAACCAGGGAGAGAUGG	sense antisense
81	MINA	CGGCUCCUGUAAGCAGAUGAAGUU AACUUCAUCUGCUUACAGGGAGCCG	sense antisense
82	NPM3	UUACACUCGUCUUUGGCUCCUCGG CCGAGGGAGCCAAAGACGAGUGUAA	sense antisense
83	TINP1	CAGAGAGGGACAAUCUCGAGCUAAA UUUAGCUCGAGAUUGUCCUCUCUG	sense antisense
84	FRG1	UAGACUUCACGUAGGAGUACUCGGC GCCGAGUACUCCUACGUGAAGUCUA	sense antisense
85	PELP1	CCUCAAUUCCUGGAGCAUCGGUAGA UCUACCGAUGCUC CAGGAAUUGAGG	sense antisense
86	ZFP106	GGUGGCAUGGUUGCUCUCUGAUUUU AAUAUCAGAGAGCAACCAUGCCACC	sense antisense
87	PWP1	CAGUGCAAUUUAGAGGUGCAUGUUU AAACAUGCACCUCUAAAUUGCACUG	sense antisense
88	PPAN	AACUGUGGAACAUCACUUUGCCCUC GAGGGCAAAGUGAUGUCCACAGUU	sense antisense

89	LAS1L	CCUACAUCCUCAGAUGGACCGUUGA UCAACGGUCCAUCUGAGGAUGUAGG	sense antisense
90	SRFBP1	GAGCAAUUGCCAGACUAGCAGUACA UGUACUGCUAGUCUGGCAAUUGCUC	sense antisense
91	NAT10	GCGGUGGCAUUUUGGGUACUCCAAUA UAUUGGAGUACCCAAAUGCCACCGC	sense antisense
92	DUSP11	UACAGUGGACACCAAUAAGUUUAUC GAUAAACUUAUUGGUGUCCACUGUA	sense antisense
93	EBNA1BP2	CCCUACGAAGCGACCCACUGAUUAU AUAUACAGUGGGUCGCUUCGUAGGG	sense antisense
94	RPL26	AAUCCCUUUGUGACUUCGACCGAA UUCGGUCGGAAGUCACAAAGGGAUU	sense antisense
95	GNL3	ACAACUUGCAUGCUCUCCUUGUAAGCC GGCUUACAAGGAGCAUGCAAGUUGU	sense antisense
96	PES1	CCAUUGUCAACAAGUUCGUGAAUA UAUUCACGGAACUUGUUGACAAUGG	sense antisense
97	NOM1	GGAUCAGCAGGAGAGAGAAAUCAUU AAUGAUUUCUCUCUCCUGCUGAUCC	sense antisense
98	NOL1	GGUCAAGACUGGACUAGUGGUGUA UACACCACUAGUCCAGUCUUUGACC	sense antisense
99	C14orf169	CAUGGAACUGUUGCUUGGUUCUUAU AUAAGAACCAAGCAACAGUCCAUG	sense antisense
100	RRP1B	CAGGAGGUUUCAGUCAGGAAGAACU AGUUCUUCUGACUGAAACCUCCUG	sense antisense
101	RPL5	UUCACUUGGUAUCUCUUAAGUAGG CCUACUUUAAGAGAUACCAAGUGAA	sense antisense
102	RPL11	GCUAGAUACACUGUCAGAUCUUCUUG CAAAGGAUCUGACAGUGUAUCUAGC	sense antisense
103	RPL23	CAGCAGUGGUCAUUCGACAACGAAA UUUCGUUGUCGAAUGACCACUGCUG	sense antisense
104	NPM1	UAGUAGCGGUGGUCAGACAUGGAAA UUUCCAUGUCUGACCACCGCUACUA	sense antisense
105	NCL	UUCUGUUGCACUGUAGGAGAGGUUG CAACCUCUCCUACAGUGCAACAGAA	sense antisense
106	NS	ACAACUUGCAUGCUCUCCUUGUAAGCC GGCUUACAAGGAGCAUGCAAGUUGU	sense antisense

107	ARF	GAAAGAACCAGAGAGGCUCUGAGAA UUCUCAGAGCCUCUCUGGUUCUUUC	sense antisense
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