

Supporting Information

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APAF1-Binding Long Noncoding RNA Promotes Tumor Growth and Multidrug Resistance in Gastric Cancer by Blocking Apoptosome Assembly

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Supporting Information

APAF1-binding long non-coding RNA promotes tumor growth and multidrug resistance in gastric cancer by blocking apoptosome assembly

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This PDF file includes:

Figure S1. ABL expression is elevated in GC.

Figure S2. Identification of the ABL-binding protein.

Figure S3. ABL promotes GC cell survival and multidrug resistance *in vitro*.

Figure S4. Knockdown of ABL promotes GC cell apoptosis by competitively blocking the APAF1 interaction with Cyt c.

Figure S5. IGF2BP1 binds and recognizes the METTL3-mediated m⁶A modification on ABL, maintaining ABL stability.

Figure S6. Identification of the effect of ABL-specific siRNA-loaded PEG-CLs.

Figure S7. ABL-specific siRNA-loaded PEG-CLs have no obvious systemic toxicity.

Table S1. Representative up or down-regulated lncRNAs from RNA-seq.

Table S2. Biotinylated ABL sense pull-down followed by MS.

Table S3. Particle size and potential of siRNA/protamine complex.

Table S4. Particle size and potential of siRNA/CLs.

Table S5. Particle size and potential of siRNA/PEG-CLs.

Table S6. The sequences of siRNAs.

Table S7. Antibodies for WB, RIP, IF, IP, and IHC.

Table S8. The oligonucleotides were used in this study.

Supporting Figures

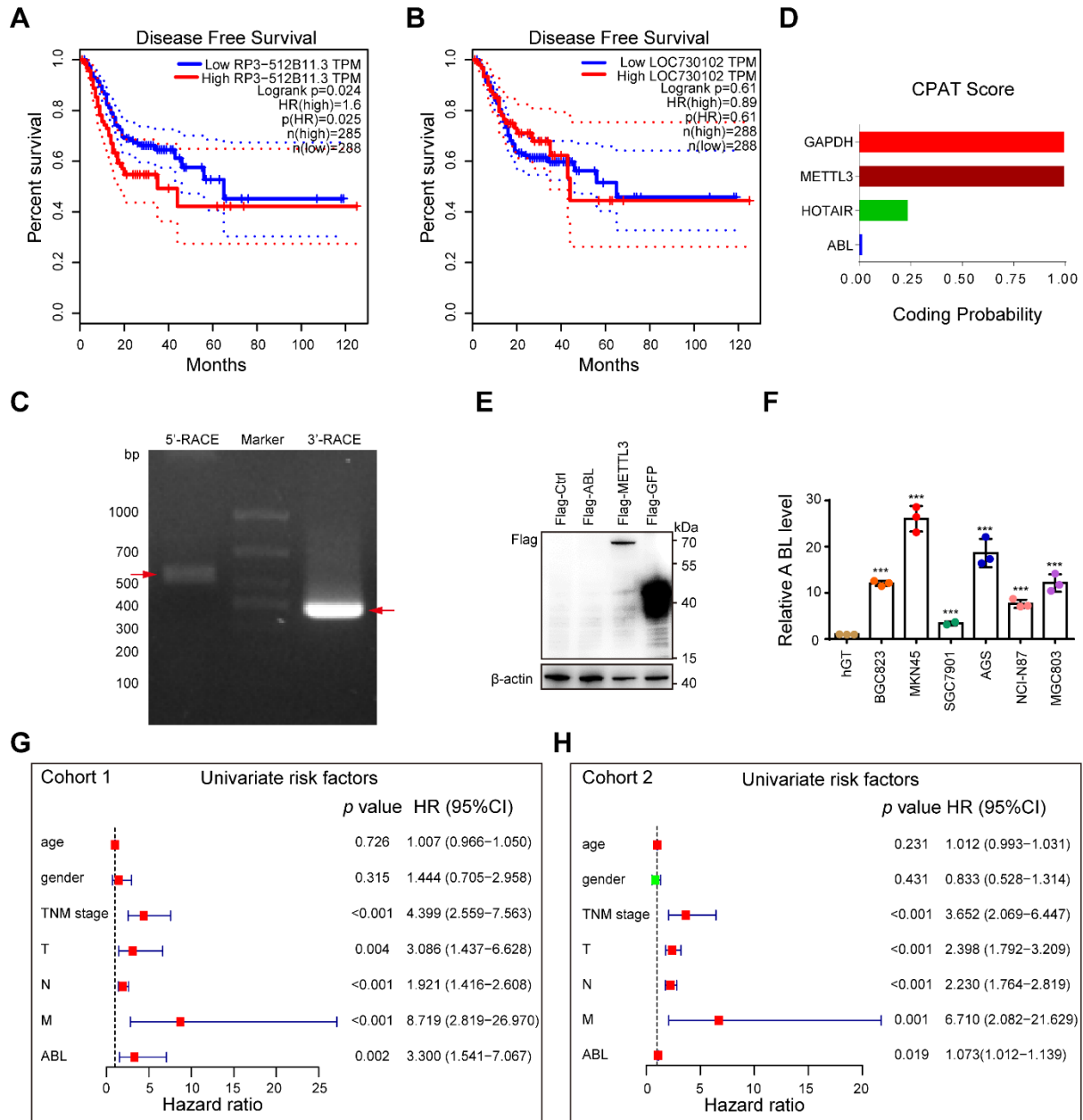


Fig. S1

Figure S1. ABL expression is elevated in GC. A-B) Kaplan-Meier disease-free survival curves based on ABL (A) or LOC730102 (B) expression using the online bioinformatics tool GEPIA (<http://gepia.cancer-pku.cn/>). C) Representative image of PCR products from the 5'- and 3'-RACE assay. D) The coding capacity of ABL, HOTAIR, METTL3, and GAPDH was determined by the CPAT webserver (<http://lilab.research.bcm.edu/cpat/>). E) HEK293T cells were transfected with the pcDNA3.1-Flag vector, ABL-Flag (full-length ABL sequence inserted

into the pcDNA3.1 vector), GFP-Flag, or METTL3-Flag. Total cell lysates were resolved in SDS-PAGE gel, and Western blotting for Flag and GAPDH was performed; GFP-Flag and METTL3-Flag were the coding proteins, which were used as positive controls. F) ABL expression was detected in human gastric mucosa tissue (hGT) and different GC cell lines by qRT-PCR. G-H) Univariate analyses were performed for GC cohort 1 (G) and cohort 2 (H). All the bars correspond to 95% confidence intervals (CIs); AUC, area under the curve; CI, confidence interval; GC, gastric cancer; HR, hazard ratio; TNM, tumor, node, metastasis. The probability of differences in DFS was ascertained by the Kaplan-Meier method with the log-rank test (A and B). The data were analyzed by a two-tailed unpaired Student's t-test (F). The data are represented as the means \pm SEM. * $p < 0.05$; ** $p < 0.01$; *** $p < 0.001$, NS, no significance.

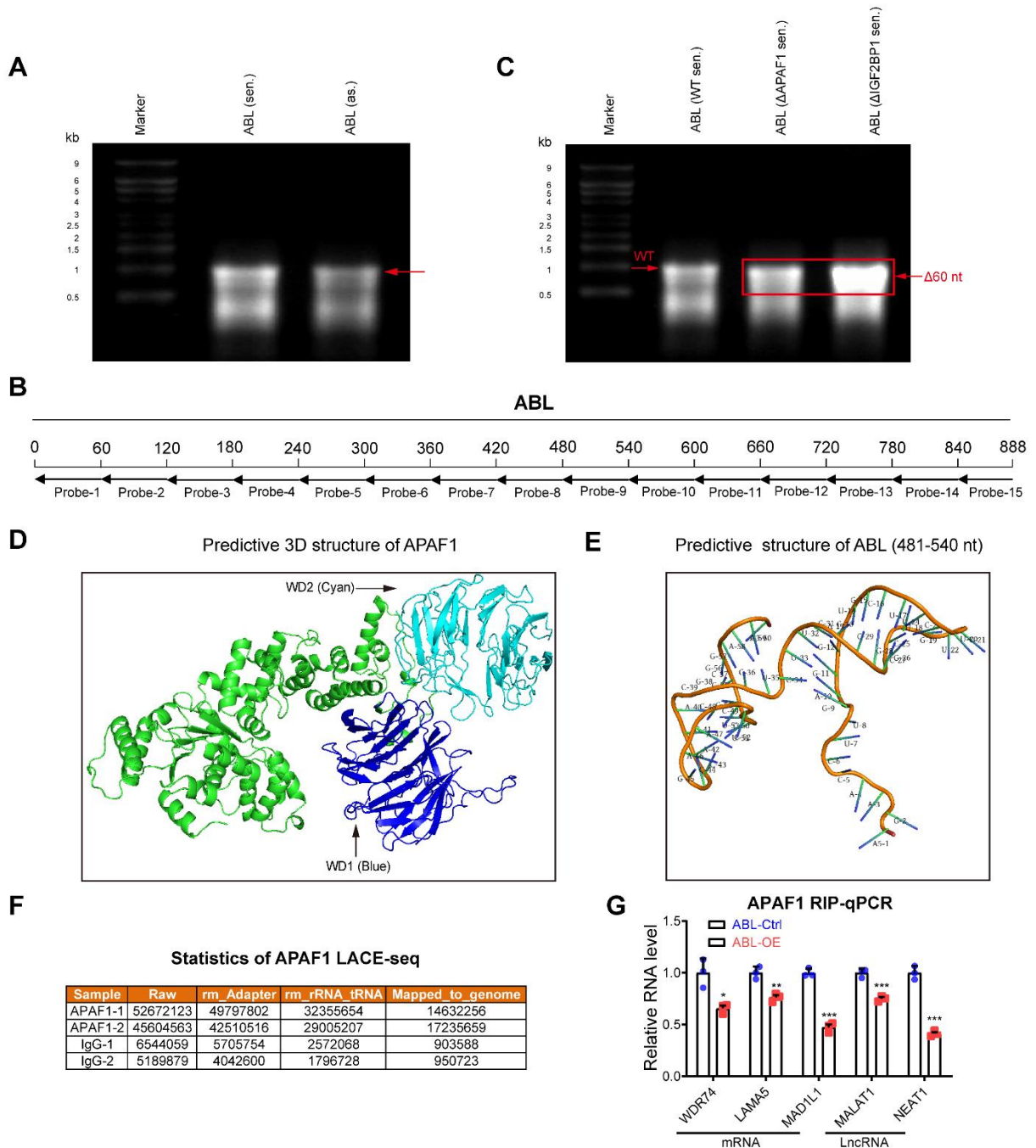


Fig. S2

Figure S2. Identification of the ABL-binding protein. A) Sense and antisense ABL transcripts were prepared *in vitro* and identified. B) Schematic diagram of ABL probes used in dot blot assay. C) Transcripts for full-length sense ABL, ABL with deletion of nt 481-540, and ABL with deletion of nt 121-160 were prepared *in vitro* and identified. D) Predicted structure of APAF1. E)

Predicted structure of ABL (nt 481-540). F) The number of reads for the APAF1 LACE-seq library. rm_Adapter, after trimming adapters; rm_rRNA_tRNA, after removing rRNA and tRNA-derived reads; Mapped_to_genome, reads mapped to the reference genome. (G) RIP-qPCR analysis was employed to detect the interaction between APAF1 and indicated mRNAs or lncRNAs upon overexpression of ABL. The data were analyzed by a two-tailed unpaired Student's t-test (G). The data are represented as the means \pm SEM. * $p < 0.05$; ** $p < 0.01$; *** $p < 0.001$, NS, no significance.

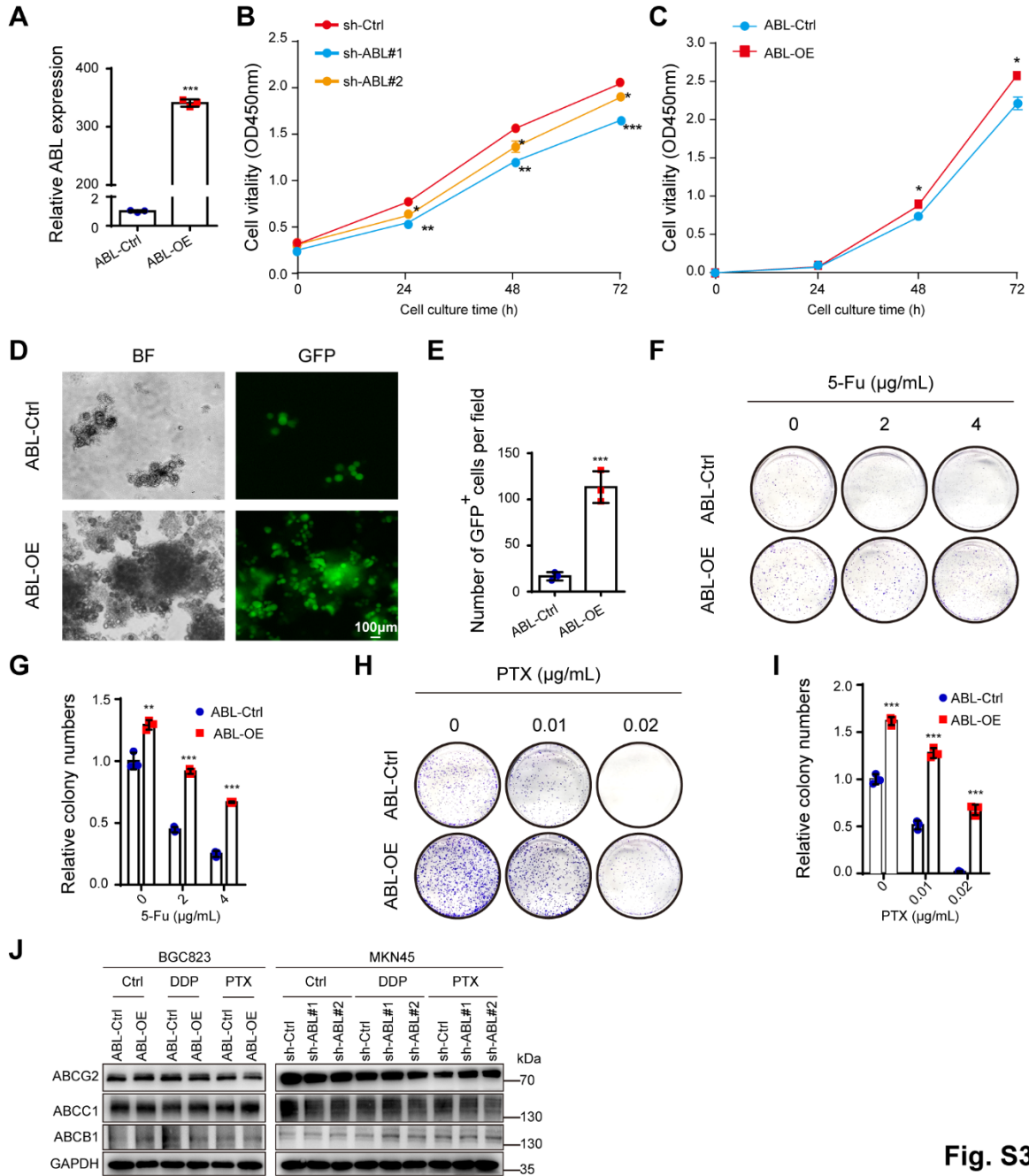


Fig. S3

Figure S3. ABL promotes GC cell survival and multidrug resistance *in vitro*. A) The expression of ABL was examined in ABL-overexpressing BGC823 cells by qRT-PCR. B-C) CCK-8 assays were used to determine the viability of ABL-deficient MKN45 cells (B) and ABL-overexpressing BGC823 cells (C). D) Suspension growth assays were used to determine the cell survival of ABL-overexpressing BGC823 cells (scale bars=100 µm). E) Quantification of the GFP⁺ cells in (D). F) Overexpression of ABL increased the colony-forming ability of

BGC823 cells and antagonized 5-Fu-induced apoptosis in these cells. G) Quantification of the colony formation assay results in (F). H) Overexpression of ABL increased the colony-forming ability of BGC823 cells and antagonized PTX-induced apoptosis in these cells. I) Quantification of the colony formation assay results in (H). J) Western blotting was applied to determine the expression of ABCB1, ABCC1 and ABCG2 in ABL-overexpressing BGC823 cells or ABL-deficient MKN45 cells after DDP (1 $\mu\text{g/ml}$) or PTX treatment (0.05 $\mu\text{g/ml}$) for 24 h. The data were analyzed by a two-tailed unpaired Student's t-test (A-C, E, G, and I). The data are represented as the means \pm SEM. * $p < 0.05$; ** $p < 0.01$; *** $p < 0.001$, NS, no significance.

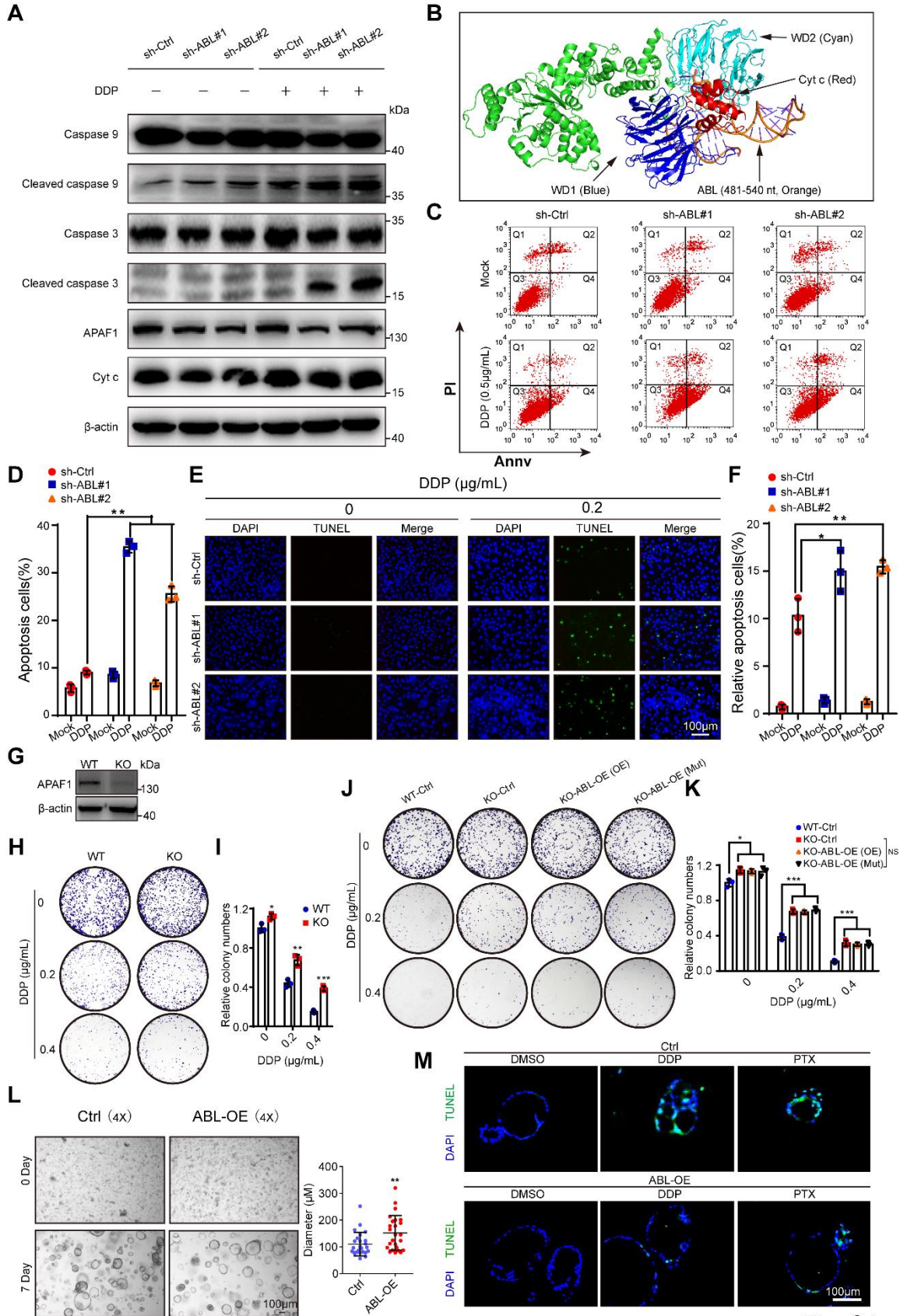


Fig. S4

Figure S4. Knockdown of ABL promotes GC cell apoptosis by competitively blocking the APAF1 interaction with Cyt c. A) Western blotting was applied to determine the expression of the indicated proteins in ABL-deficient MKN45 cells after DDP treatment at 0 or 1 $\mu\text{g/ml}$ for 24 h. B) Docking analysis of protein-protein and RNA-protein interaction models for APAF1 and ABL (nt 481–540) and APAF1-Cyt c. C) An annexin-V-FITC/PI assay was used to detect apoptotic ABL-deficient MKN45 cells after DDP treatment at 0 or 1 $\mu\text{g/ml}$ for 24 h. D) Quantification of the apoptotic cells in (C). E) A TUNEL assay was used to detect apoptotic ABL-deficient MKN45 cells after DDP treatment at 0 or 1 $\mu\text{g/ml}$ for 24 h. F) Quantification of the apoptotic cells in (E). G) The protein levels of APAF1 in BGC823 cells with APAF1 knockout were measured by western blotting. H) Knockout of APAF1 increased the colony-forming ability of BGC823 cells and antagonized DDP-induced apoptosis in these cells. I) Quantification of the colony formation assay results in (H). J) The colony-forming ability was detected in APAF1 knockout BGC823 cells transfected with ABL-overexpressing WT, Mut plasmids, or their corresponding controls after DDP treatment at 0, 0.2, 0.4 $\mu\text{g/ml}$ for 24 h. K) Quantification of the colony formation assay results in (J). L) Representative images of GC organoids transfected with ABL overexpression vectors or a control lentivirus at 0 or 7 days (scale bars=100 μm , left panel, n=3) and the quantification of organoid diameters (right panel). M) Sections of organoids from different groups underwent TUNEL staining (scale bars=100 μm). The data were analyzed by a two-tailed unpaired Student's t-test (D, F, I, K, and L). The data are represented as the means \pm SEM. * $p<0.05$; ** $p<0.01$; *** $p<0.001$, NS, no significance.

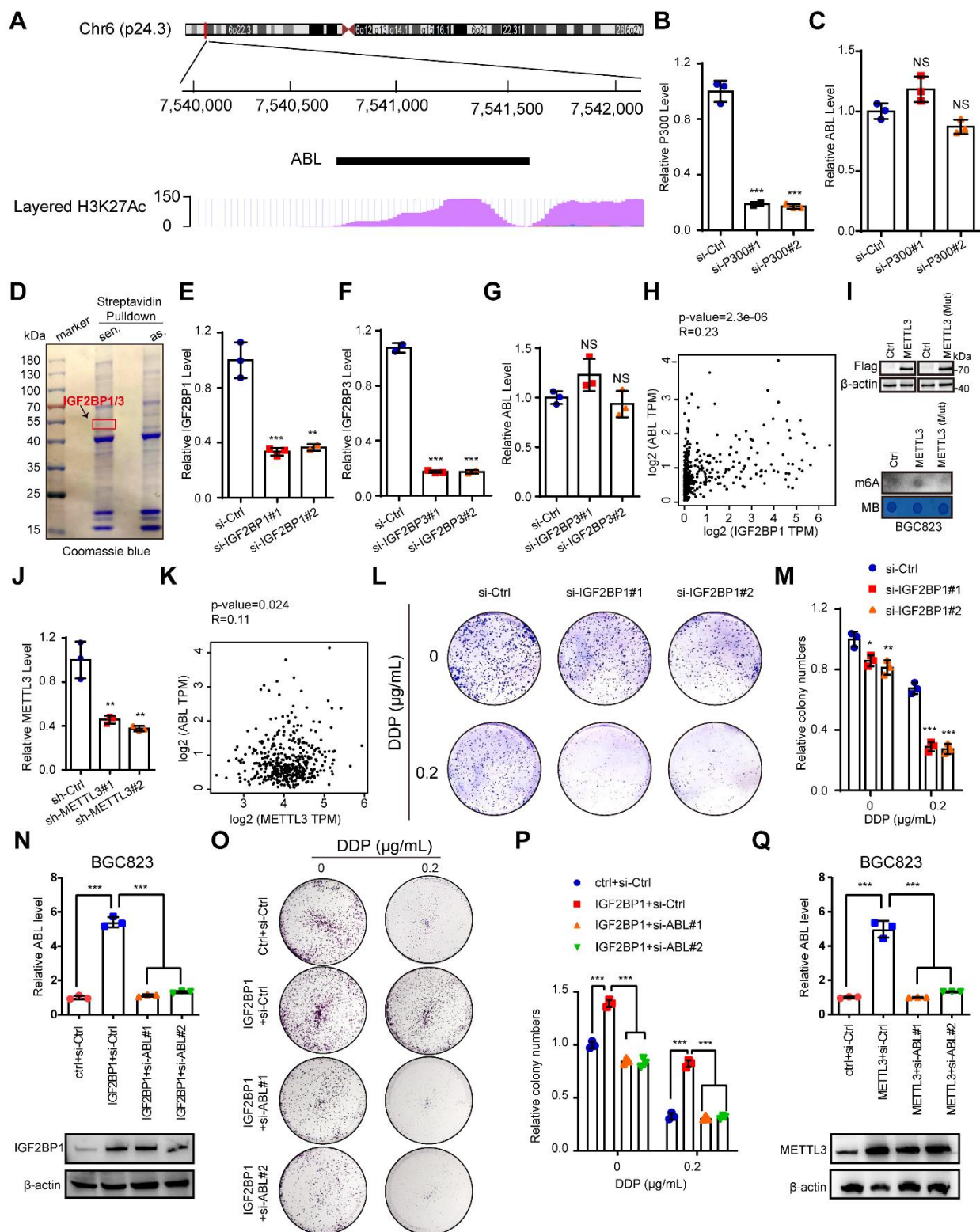


Fig.S5

Figure S5. IGF2BP1 binds and recognizes the METTL3-mediated m6A modification on ABL, maintaining ABL stability. A) Data from the UCSC genome bioinformatics site

(<http://genome.ucsc.edu/>) showed enrichment of H3K27ac in the promoter of ABL. B) P300 knockdown efficiency was verified in BGC823 cells by qRT-PCR. C) The ABL levels in BGC823 cells with P300 deficiency were determined by qRT-PCR. D) Coomassie brilliant blue staining of proteins pulled down by biotinylated ABL. E-F) IGF2BP1 and IGF2BP3 knockdown efficiencies in BGC823 cells were verified by qRT-PCR. G) The ABL levels in BGC823 cells with IGF2BP3 deficiency were determined by qRT-PCR. H) IGF2BP1 expression was positively correlated with ABL expression in GC (linear regression) using the online bioinformatics tool GEPIA (<http://gepia.cancer-pku.cn/>). TPM, transcripts per million. I) The protein levels of Flag-METTL3 in BGC823 cells with wild-type or catalytic mutant (Mut) METTL3 overexpression were measured by western blotting (upper panel). The mRNAs isolated from wild-type or catalytic mutant METTL3-overexpressing GC cells were used in dot blot analyses with an m⁶A antibody (bottom panel). MB (methylene blue) staining served as a loading control. J) The knockdown efficiencies of METTL3 were verified in MKN45 GC cells by qRT-PCR. K) METTL3 expression was positively correlated with ABL expression in GC (linear regression) using the online bioinformatics tool GEPIA (<http://gepia.cancer-pku.cn/>). L) Knockdown of ABL decreased the colony-forming ability of BGC823 cells and promoted DDP-induced apoptosis in these cells. M) Quantification of the colony formation assay results in (L). N) The knockdown efficiency of ABL using specific siRNAs in IGF2BP1-overexpressing BGC823 cells was detected by qRT-PCR (upper panel); The protein levels of IGF2BP1 in BGC823 cells with IGF2BP1 overexpression were measured by western blotting (bottom panel). O) Representative images of the cell colony formation abilities of IGF2BP1-overexpressing BGC823 cells transfected with ABL-specific siRNAs or corresponding controls and then treated with DDP at the indicated doses for 24 h. P) Quantification of the colony formation assay results in (O). Q) The knockdown efficiency of ABL using specific siRNAs in METTL3-overexpressing BGC823 cells was detected by qRT-PCR (upper panel); The protein levels of METTL3 in BGC823 cells with METTL3 overexpression were measured by western blotting (bottom panel). The data were analyzed by a two-tailed unpaired Student's t-test (B, C, E-G, J, M, N, P, and Q). The data are represented as the means \pm SEM. * p<0.05; ** p<0.01; *** p<0.001, NS, no significance.

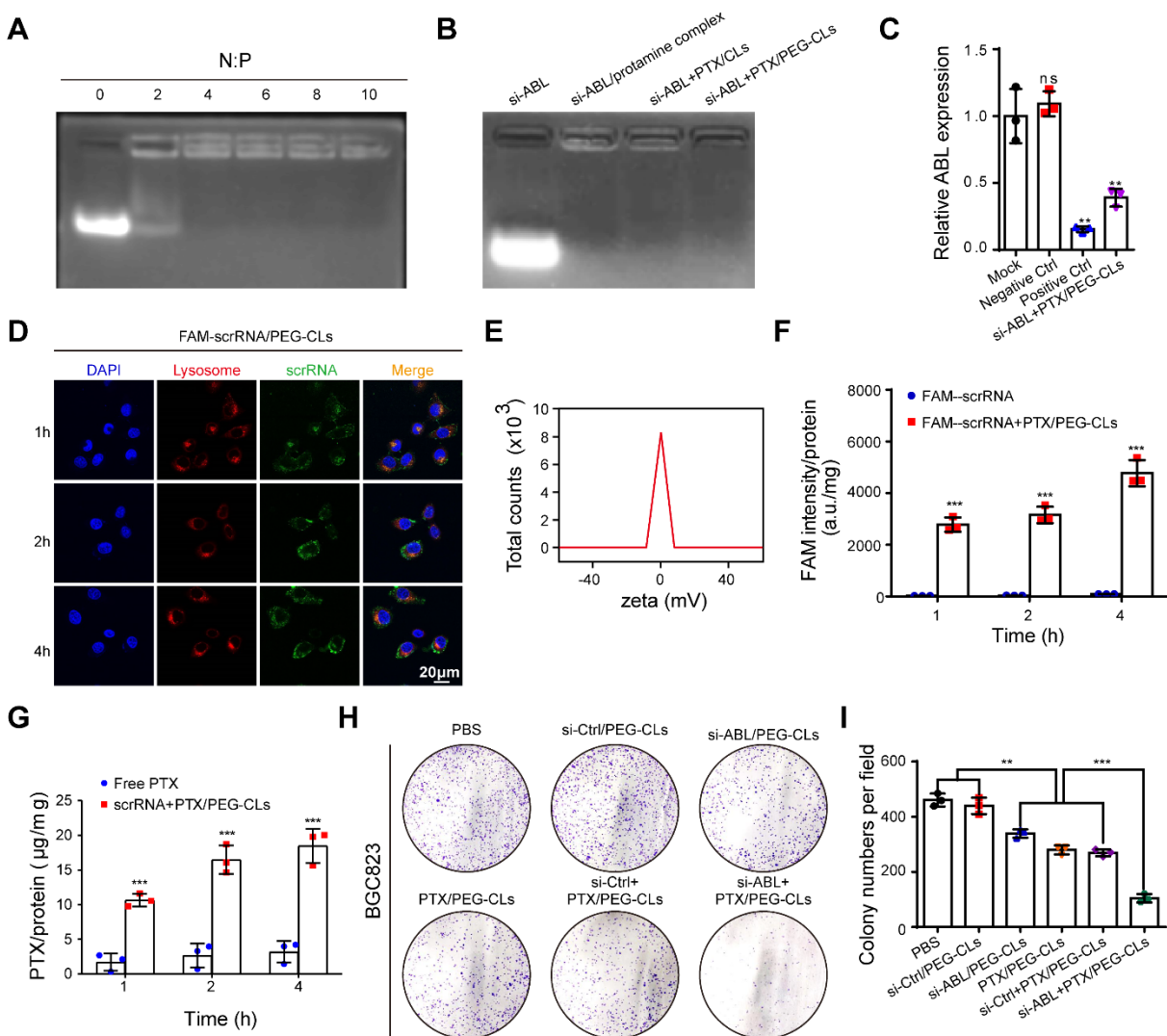


Fig. S6

Figure S6. Identification of the effect of ABL-specific siRNA-loaded PEG-CLs. A) Agarose gel electrophoresis of the siRNA/protamine complex at different N:P ratios. B) Agarose gel electrophoresis of si-ABL, si-ABL/protamine complex, si-ABL+PTX/CLs, and si-ABL+PTX/PEG-CLs. C) The postinsertion group showed no obvious effect on transfection efficiency, indicating that the postinsertion method is better for preparing PEG-CLs. D) Intracellular delivery of siRNA PEG-CLs or free siRNA was detected at the indicated times by IF (scale bars=20 μm). E) Zeta potential of ABL-specific siRNA-loaded PEG-CLs. F-G) The cellular uptake of free siRNA, PTX, and siRNA+PTX/PEG-CLs was detected. H) A colony formation assay was used to assess the *in vitro* antitumor capability of si-ABL+PTX/PEG-CLs. I) Quantification of the colony formation assay results in (H). The data were analyzed by a

two-tailed unpaired Student's t-test (C, F, G, and I). The data are represented as the means \pm SEM. * $p < 0.05$; ** $p < 0.01$; *** $p < 0.001$, NS, no significance.

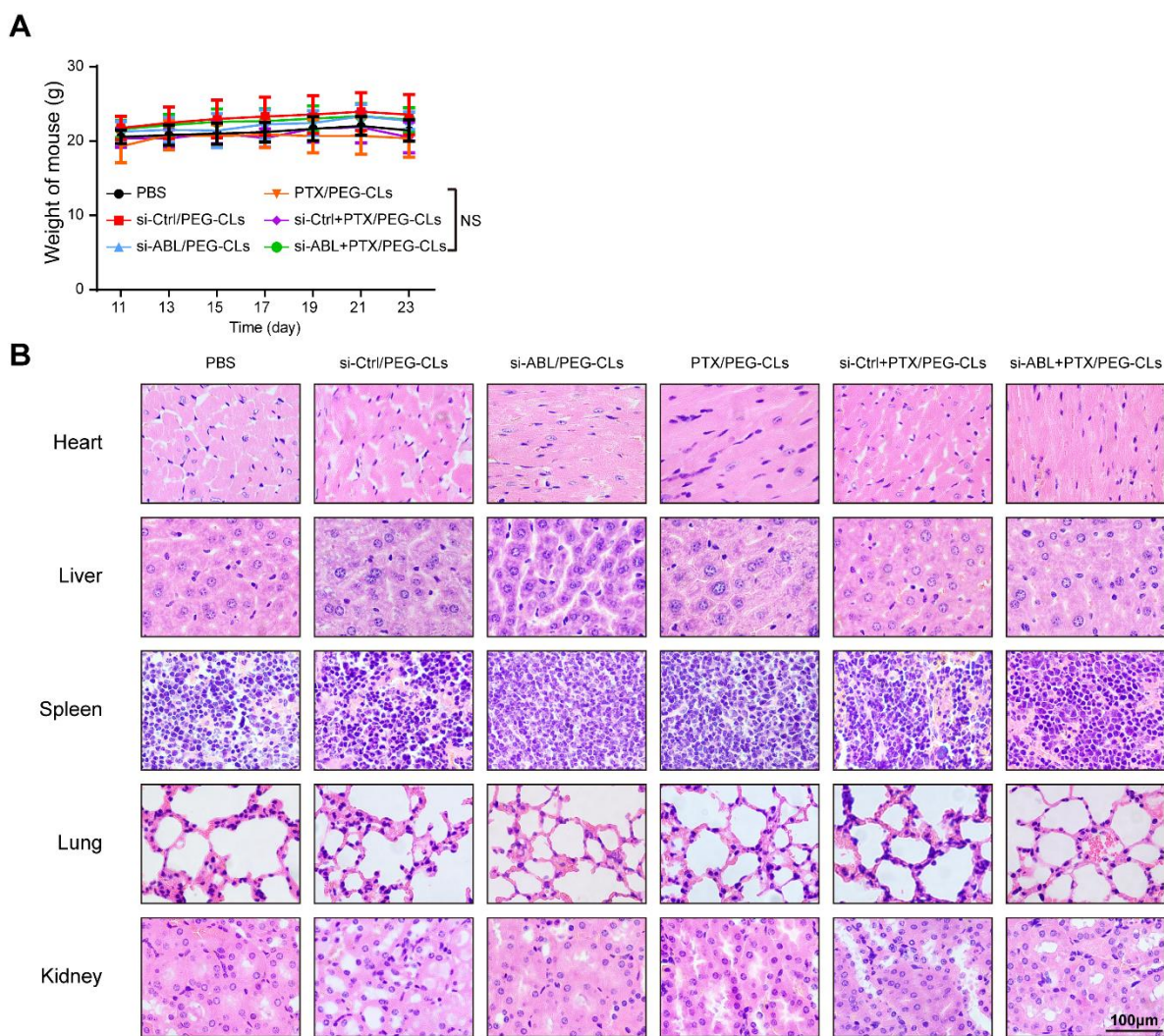


Fig. S7

Figure S7. ABL-specific siRNA-loaded PEG-CLs have no obvious systemic toxicity. A) Mouse weight was monitored every other day in each group (n=6). B) Representative H&E staining of major organs, including the heart, liver, spleen, lungs, and kidneys, from mice in each group at the end of the experiment. The data were analyzed by one-way ANOVA test followed by Turkey's multiple comparisons (A). The data are represented as the means \pm SEM. * $p < 0.05$; ** $p < 0.01$; *** $p < 0.001$, NS, no significance.

Supporting Tables

Table S1. Representative up or down-regulated lncRNAs from RNA-seq

LncRNA Annotation		Differential Expression			
GeneSymbol	source	log2FC	Fold_Change	p_value	q_value
RP3-512B11.3	GENCODE	2.348886063	5.094307562	1.40887E-05	0.039354 458
LOC730102	RefSeq	1.990616039	3.974066569	1.8763E-05	0.043010 915
RP11-37B2.1	GENCODE	1.305542559	2.471766665	0.001497517	0.158735 494
LOC100288637	RefSeq	1.162140427	2.23789202	0.001645347	0.162808 241
RP11-1275H24.3	GENCODE	1.167614434	2.246399367	0.001821883	0.165664 29
DUXAP10	RefSeq	2.675223575	6.387376868	0.002724512	0.185397 161
LINC00543	GENCODE	1.282344685	2.432339625	0.004594469	0.203758 381
RP11-803D5.4	GENCODE	1.358719515	2.56457456	0.004865512	0.207919 361
RP11-649A18.4	GENCODE	1.570332954	2.969732436	0.008310237	0.236849 722
LINC01296	RefSeq	1.725211187	3.306285223	0.01347973	0.267398 003
MIR4435-2HG	RefSeq	1.038804638	2.054524648	0.015748694	0.277687 696
RP11-589N15.2	GENCODE	1.217501639	2.325436652	0.016904124	0.281968 239
CTA-384D8.34	GENCODE	1.17969342	2.265286334	0.017907056	0.286048 666
LOC401585	RefSeq	2.345952443	5.083959168	0.019422357	0.291005

					463
LINC01572	RefSeq	1.048726356	2.068702744	0.026202525	0.310253
					643
RP11-649A18.5	GENCODE	1.081856016	2.116757523	0.027562718	0.313953
					485
ABHD11-AS1	RefSeq	1.83811065	3.57541486	0.02943696	0.317886
					269
LINC00152	RefSeq	1.366649814	2.578710494	0.030982672	0.320545
					784
RP11-79H23.3	GENCODE	1.324402887	2.504292168	0.033006479	0.324641
					191
LL22NC03-N14H	GENCODE	1.672258134	3.187130592	0.036003641	0.332974
11.1					077
RP11-797A18.6	GENCODE	1.428063306	2.69085249	0.037935589	0.337272
					614
AC017002.2	GENCODE	1.281113686	2.430265083	0.038046265	0.337272
					614
MIR4435-2HG	RefSeq	1.232520427	2.349771429	0.040180489	0.342033
					401
GSE61474_XLOC	RNA-seq:	1.210444469	2.314089188	0.040458719	0.343015
_058624	Clark et al				034
	2015				
LINC00857	RefSeq	1.058650108	2.082981619	0.04302053	0.348952
					716
RP11-37B2.1	GENCODE	1.492443998	2.813652186	0.043948088	0.350914
					696
RP11-626H12.1	GENCODE	2.317480806	4.984610605	0.0443468	0.352218
					925
LOC400043	RefSeq	-2.21287222	0.21570444	0.043595338	0.349999
					719
PWAR5	RefSeq	-2.03395636	0.244184518	0.024637721	0.308751

PWAR6	GENCODE	-1.76530452	0.294164586	0.019661114	0.291746
					826
GSE61474_XLOC_020344	RNA-seq: Clark et al 2015	-1.734616	0.300488983	0.030932554	0.320374
					778
PART1	RefSeq	-1.70357706	0.307023914	0.006058087	0.221302
					4
MIR4458HG	RefSeq	-1.07645654	0.474192075	0.025287889	0.309135
					684
LINC01184	RefSeq	-1.04697879	0.483980628	0.032289326	0.323317
					663

Table S2. Biotinylated ABL sense pull-down followed by MS

Band 1 for peptides identification		S	C	P	Uni	P	P	A	M	c
Acc	Description	c	ov	ro	que	ep	S	A	W	al
on		o	er	te	Pept	ti	M	s	[k	c.
		r	ag	in	ides	de	s		Da	p
		e	e	s		s] I	
O14	Isoform 3 of Apoptotic protease-activating factor 1 OS=Homo sapiens OX=9606 GN=APAF1 - [APAF_HUMAN]	6	65	8	70	70	2	1	13	6.
727-		1	.4				1	1	5.9	4
3		3.	9				6	9		4
		4						4		
		6								
Q02	Desmoglein-1 OS=Homo sapiens OX=9606 GN=DSG1 PE=1 SV=2 - [DSG1_HUMAN]	2.	1.	1	1	1	1	1	11	5.
413		8	53					0	3.7	0
		3						4		3
								9		
Q14	Desmoglein-2 OS=Homo sapiens OX=9606	2.	5.	1	4	4	5	1	12	5.

126	GN=DSG2 PE=1 SV=2 - [DSG2_HUMAN]	8	19					1	2.2	2
		4						1		4
								8		
P42	Lamina-associated polypeptide 2, isoform	1.	1.	1	1	1	1	6	75.	7.
166	alpha OS=Homo sapiens OX=9606	7	59					9	4	6
	GN=TMPO PE=1 SV=2 - 0							4		6
	[LAP2A_HUMAN]									
P08	Heat shock protein HSP 90-beta OS=Homo	1.	4.	7	2	2	2	7	83.	5.
238	sapiens OX=9606 GN=HSP90AB1 PE=1	6	01					2	2	0
	SV=4 - [HS90B_HUMAN]	9						4		3
Q9	Thyroid hormone receptor-associated protein	1.	1.	1	1	1	1	9	10	1
Y2	3 OS=Homo sapiens OX=9606 GN=THRAP3	6	15					5	8.6	0.
W1	PE=1 SV=2 - [TR150_HUMAN]	3						5		1
										5
P02	Isoform 5 of Prelamin-A/C OS=Homo	1.	5.	9	2	2	2	5	62.	6.
545-	sapiens OX=9606 GN=LMNA - 6	31						6	8	1
5	[LMNA_HUMAN]	1						5		8
Q13	Isoform 2 of Protein flightless-1 homolog	1.	0.	3	1	1	1	1	13	6.
045-	OS=Homo sapiens OX=9606 GN=FLII - 6	74						2	8.4	0
2	[FLII_HUMAN]	1						1		7
								4		
P78	Isoform 2 of General transcription factor II-I	0.	0.	4	1	1	1	9	10	7.
347-	OS=Homo sapiens OX=9606 GN=GTF2I - 0	94						5	7.9	9
2	[GTF2I_HUMAN]	0						7		4
P27	Inositol-trisphosphate 3-kinase B OS=Homo	0.	2.	1	1	1	3	9	10	8.
987	sapiens OX=9606 GN=ITPKB PE=1 SV=5 - 0	22						4	2.3	4
	[IP3KB_HUMAN]	0						6		3
Q96	Isoform 2 of Histone-lysine	0.	0.	3	1	1	1	2	26	8.
L73	N-methyltransferase, H3 lysine-36 and H4	0	58					4	7.2	5
-2	lysine-20 specific OS=Homo sapiens	0						2		1
	OX=9606 GN=NSD1 - [NSD1_HUMAN]							7		

Band 2 for peptides identification											
Q9	Insulin-like growth factor 2 mRNA-binding	1	7.	9	2	3	7	5	63.	9.	
NZI	protein 1 OS=Homo sapiens OX=9606	4.	80					7	4	2	
8	GN=IGF2BP1 PE=1 SV=2 - [IF2B1_HUMAN]	1						7		0	
A0	Serum albumin OS=Homo sapiens OX=9606	4.	3.	9	1	1	1	3	45.	6.	
A08	GN=ALB PE=1 SV=1 - [A0A087WWT3_HUMAN]	4	79					9	1	1	
7W	[A0A087WWT3_HUMAN]	0						6		0	
WT											
3											
F5G	Retinoic acid-induced protein 3 (Fragment)	2.	4.	2	1	1	1	2	30.	9.	
WG	OS=Homo sapiens OX=9606 GN=GPRC5A	4	76					7	5	0	
3	PE=1 SV=8 - [F5GWWG3_HUMAN]	1						3		3	
Q07	Isoform 3 of KH domain-containing,	2.	3.	3	1	1	1	4	44.	7.	
666-	RNA-binding, signal transduction-associated	2	47					0	0	2	
3	protein 1 OS=Homo sapiens OX=9606	7						4		8	
	GN=KHDRBS1 - [KHDR1_HUMAN]										
H0	Polyadenylate-binding protein 2 (Fragment)	2.	11	6	1	1	1	9	11.	1	
YJH	OS=Homo sapiens OX=9606 GN=PABPN1	2	.4					6	1	1.	
9	PE=1 SV=1 - [H0YJH9_HUMAN]	2	6							2	
										7	
P50	Isoform 3 of T-complex protein 1 subunit	2.	2.	3	1	1	1	4	51.	5.	
990-	theta OS=Homo sapiens OX=9606	1	11					7	6	2	
3	GN=CCT8 - [TCPQ_HUMAN]	8						5		4	
B4	Pyruvate kinase OS=Homo sapiens OX=9606	2.	23	9	7	7	7	4	49.	7.	
DN	GN=PKM PE=1 SV=1 - [B4DNK4_HUMAN]	1	.1					5	9	8	
K4	[B4DNK4_HUMAN]	4	9					7		3	
Q8I	Mitotic interactor and substrate of PLK1	2.	2.	1	1	1	1	6	75.	6.	
VT2	OS=Homo sapiens OX=9606 GN=MISP	0	36					7	3	8	
	PE=1 SV=1 - [MISP_HUMAN]	7						9		3	
P06	Isoform 3 of Tyrosine-protein kinase Fyn	2.	2.	7	1	1	2	4	54.	6.	

241-	OS=Homo sapiens	OX=9606	GN=FYN	-	0	28					8	5	2
3	[FYN_HUMAN]					4					2		3
Q15	2'-5'-oligoadenylate synthase-like protein				2.	6.	3	2	2	2	5	59.	7.
646	OS=Homo sapiens	OX=9606	GN=OASL		0	03					1	2	8
	PE=1 SV=2 - [OASL_HUMAN]					3					4		7
D6	Drebrin (Fragment)	OS=Homo sapiens			2.	6.	5	1	1	1	3	36.	5.
R9	OX=9606	GN=DBN1	PE=1	SV=1	-	0	62				1	4	1
W4	[D6R9W4_HUMAN]					1					7		4
P38	Stress-70 protein, mitochondrial	OS=Homo sapiens			1.	1.	1	1	1	1	6	73.	6.
646	sapiens OX=9606	GN=HSPA9	PE=1	SV=2	-	9	62				7	6	1
	[GRP75_HUMAN]					5					9		6
A0	Flotillin-1 (Fragment)	OS=Homo sapiens			1.	12	17	3	3	3	3	39.	6.
A14	OX=9606	GN=FLOT1	PE=1	SV=1	-	8	.3				5	8	3
0T9	[A0A140T9R1_HUMAN]					7	2				7		7
R1													
Q8	G2/mitotic-specific cyclin-B3	OS=Homo sapiens			1.	0.	1	1	1	1	1	15	6.
W	sapiens OX=9606	GN=CCNB3	PE=1	SV=2	-	7	93				3	7.8	6
WL	[CCNB3_HUMAN]					0					9		8
7											5		
P61	Actin-related protein 3	OS=Homo sapiens			1.	6.	4	2	2	2	4	47.	5.
158	OX=9606	GN=ACTR3	PE=1	SV=3	-	6	22				1	3	8
	[ARP3_HUMAN]					3					8		8
Q9	Probable ATP-dependent RNA helicase				0.	5.	1	1	1	1	5	59.	1
NU	DDX28	OS=Homo sapiens			0	19					4	5	0.
L7	GN=DDX28	PE=1	SV=2	-	0						0		4
	[DDX28_HUMAN]												2
P35	Isoform 2 of Protein DEK	OS=Homo sapiens			0.	7.	6	2	2	2	3	38.	8.
659-	OX=9606	GN=DEK	-	[DEK_HUMAN]		0	33				4	7	1
2						0					1		5
P06	Isoform MBP-1 of Alpha-enolase	OS=Homo sapiens			0.	3.	2	1	1	1	3	36.	6.
733-	sapiens	OX=9606	GN=ENO1	-	0	52					4	9	2

2	[ENOA_HUMAN]	0	1	8
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Table S3. Particle size and potential of siRNA/protamine complex

N:P (mol:mol)	2	4	6	8	10
Size (nm)	267.9±8.4	246.4±6.1	226.1±22.9	187.5±13.7	167.4±7.8
Zeta (mV)	-27.0±2.1	-18.6±0.6	-1.7±1.2	9.7±2.3	16.8±4.8

Table S4. Particle size and potential of siRNA/CLs

Lipid (nmol/μg siRNA)	50	100
Size (nm)	146.5 \pm 5.8	68.5 \pm 1.7
Zeta (mV)	27.2 \pm 10.7	34.3 \pm 3.1

Table S5. Particle size and potential of siRNA/PEG-CLs

$n_{\text{PEG}}/n_{\text{total lipids}}$	1%	3%	5%
Size (nm)	68.9±2.1	71.0±1.0	73.1±1.9
Zeta (mV)	27.7±4.6	10.0±0.3	0.17±0.61

Table S6. The sequences of siRNAs

siRNAs	Sequences (5'-3')
ABLsiRNA#1	GCGAAGAACCCTAGGCAGA
ABLsiRNA#2	ACCCTCTCCGGA ACTCAGA
P300 siRNA#1	CGACTTACCAGATGAATTA
P300 siRNA#2	GCACAAATGTCTAGTTCTT
METTL3 siRNA#1	CGACTACAGTAGCTGCCTT
METTL3 siRNA#2	CTGCAAGTATGTTCACTATGA
IGF2BP1 siRNA#1	GGCTCAGTATGGTACAGTA
IGF2BP1 siRNA#2	TGAAGATCCTGGCCCATAA
IGF2BP3 siRNA#1	GCTGAGAAGTCGATTACTA
IGF2BP3 siRNA#2	TAAGGAAGCTCAAGATATA

Table S7. Antibodies for WB, RIP, IF, IP, and IHC

Antibodies	Source	Identifier
anti-GAPDH (for WB)	Beyotime	Cat#AG019
anti-APAF1 (for WB, RIP and LACE)	Abcam	Cat#ab234436
anti-APAF1 (for IF and IP)	Santa	Cat#sc-135836
anti-IGF2BP1 (for WB, RIP and IF)	Proteintech	Cat#22803-1-AP
anti-METTL3 (for WB)	Proteintech	Cat# 15073-1-AP
anti-Rabbit IgG (for RIP and LACE)	Beyotime	Cat#A7016
anti-Mouse IgG (for RIP)	Beyotime	Cat#A7028
anti-HRP mouse (for WB)	Beyotime	Cat#A0216
anti-HRP rabbit (for WB)	Beyotime	Cat#A0208
anti-IHC rabbit (for IHC)	Servicebio	Cat#G1215
anti- β -actin (for WB)	Beyotime	Cat#AA128
anti-caspase-9 (for WB)	CST	Cat#9509
anti-caspase-3 (for WB)	CST	Cat#9662
anti-cleaved caspase-3 (for IF)	CST	Cat#9664
anti-Cyt c (for WB and IF)	Abcam	Cat#ab133504
anti-Ki-67 (for IHC)	Servicebio	Cat#GB111499
anti-HA (for WB)	Beyotime	Cat#AF5057
anti-His (for WB)	Beyotime	Cat#P2233
anti-FLAG (for WB)	Sigma -Aldrich	Cat#F1804
anti-m6A (for RIP and Dot blot)	Abcam	Cat#ab208577
anti-GST (for WB)	Abmart	Cat#MY1901
anti-His (for WB)	SMART	Cat# SLAB28

Table S8. The oligonucleotides were used in this study

Names	Sequences (5'-3')
qPCR primers for gene expression	
ABL F ^a	TGAGGATGCTTGTCTCTCGC
ABL R ^b	GAATGTTCAAGCTCCGCGTC
GAPDH F	CATGTGGGCCATGAGGTCCACCAC
GAPDH R	GGGAAGCTCACTGGCATGGCCTTCC
METTL3 F	ATCCCCAAGGCTTCAACCAG
METTL3 R	AGGGTGATCCAGTTGGGTTG
P300 F	GCAGTGTGCCAAACCAGATG
P300 R	CATAGCCCATAGGCGGGTTG
IGF2BP1 F	AGCTCCTTTATGCAGGCTCC
IGF2BP1 R	CCGGGAGAGCTGTTTGATGT
IGF2BP3 F	ACTGCACGGGAAACCCATAG
IGF2BP3 R	CCAGCACCTCCCCTGTAAAT
WDR74 F	CCTGGGGTGTGTAGGATGC
WDR74 R	CAAGTCCAGCCAGTCATTCCG
LAMA5 F	GACTGCCAACAGTGCCAAC
LAMA5 R	CCACCCTGATAGGTGCCAT
MAD1L1 F	TGGACTGGATATTTCTACCTCGG
MAD1L1 R	CCTCACGCTCGTAGTTCCTG
MALAT1 F	AAAGCAAGGTCTCCCCACAAG
MALAT1 R	GGTCTGTGCTAGATCAAAAGGCA
NEAT1 F	CCAGTTTTCCGAGAACCAAA
NEAT1 R	ATGCTGATCTGCTGCGTATG
SCARNA2 F	GATCTTATTTGATCGGATCGTG
SCARNA2 R	CAATTCATCACTTCTGAGCGC
ABL probes used in <i>in vitro</i> RNA pull-down coupled with dot-blot assay	
Probe-1	CTATTTCCGCCCCGGGGACCGAGGAAATCCTCAAAAGCCG GGCCACGCATTCCCAGCCCCGA

	GATCAACAAAAGTAAATATTTGACCAGTAAGTCTCCAAT
Probe-2	CGAGAGGCCCTCCATTCCAG
	TCTAAAGCGTTCCTCCTCACTGTGGTCCTAGTCAATGCG
Probe-3	TGGTAGGTAGCGTTTCCAGT
	ATTCCTGAATTCCTGGAATAAAACAAAACAAAAAAG
Probe-4	CCCCAAACAAAACCTCTATCA
	GGAAGTCGTGGTTTTCTGTAAAAAAGGTTTTACACATC
Probe-5	AGAAATCGGACTTCTGGGGCA
	AGCTGCAGTGTTTTCTCTTCATGCCAAAGAGATTAATAT
Probe-6	TTACGGTGAAGACTTTTCCA
	ATCGAAAGGGAAAAATAAAAGAAATTGCAGTTTATATT
Probe-7	TTTGTTATCCCATGAAAAATT
	GGTACAAAATTTGGAAAATGGACCTGGCGTTGAAATCTT
Probe-8	TACTCCTCGTTTAAAAAGACC
	GTTCCGGAGAGGGTTCGGTCTGCGGAGAAGCGCGCGAGA
Probe-9	GACAAGCATCCTCAAGGTTCT
	GCCACCCTCAGGCTCAAGTCCCAGGAGGCCTCCTCCGCT
Probe-10	GACTTGGGTGAACAGCTCTGA
	ATGTTCAAGCTCCGCGTCACTCATCGGTACCAAGTTGATG
Probe-11	ATCTCTGCCTAGGGTTCTTC
	GCCTCTTGCAGTTAACTCAGCCTCTTAACTGGGAGTGG
Probe-12	CCCTGGAGCAAGTCGCTGGA
	GACTGGGTCTGTATCCAAACCCGGTTCGGTTCCTGGAGC
Probe-13	CAACGGCGTAGTTAACTCA
	TTACTAAAGCGTCATTTAACAATTTTTCATATTATAGAAA
Probe-14	CAGCTTTTCATGGATGGTGT
	TTGTTTGGAAAGATGAAAAATTTATTAGAAAAGATCTTT
Probe-15	AAATGGCTT

a: F, Forward; b: R, Reverse.