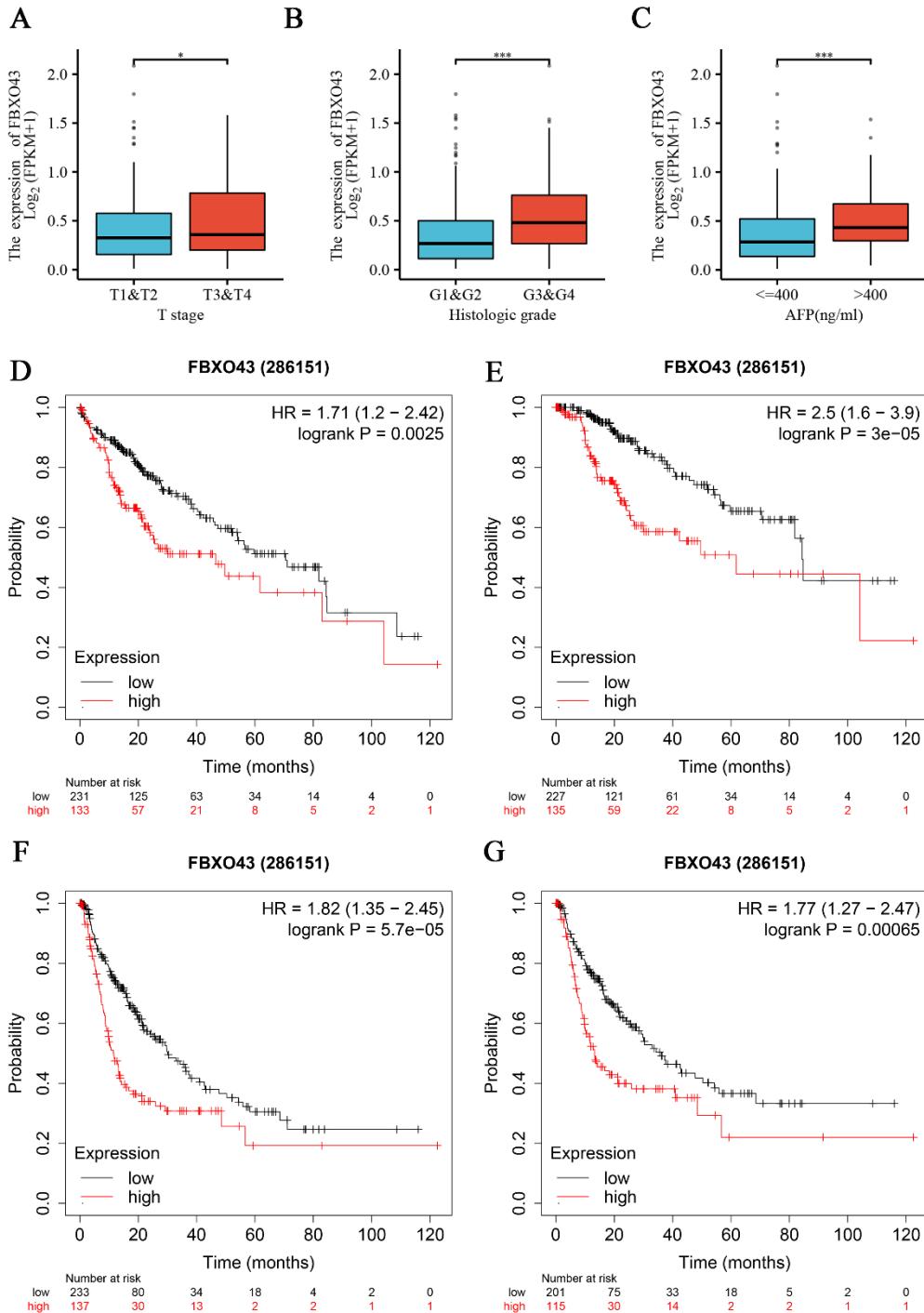
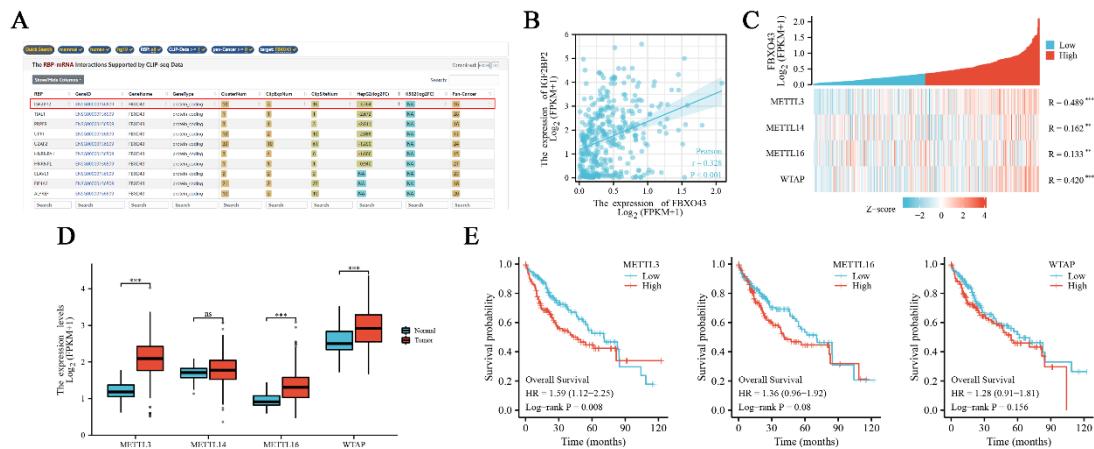


## Supplementary Figure legends

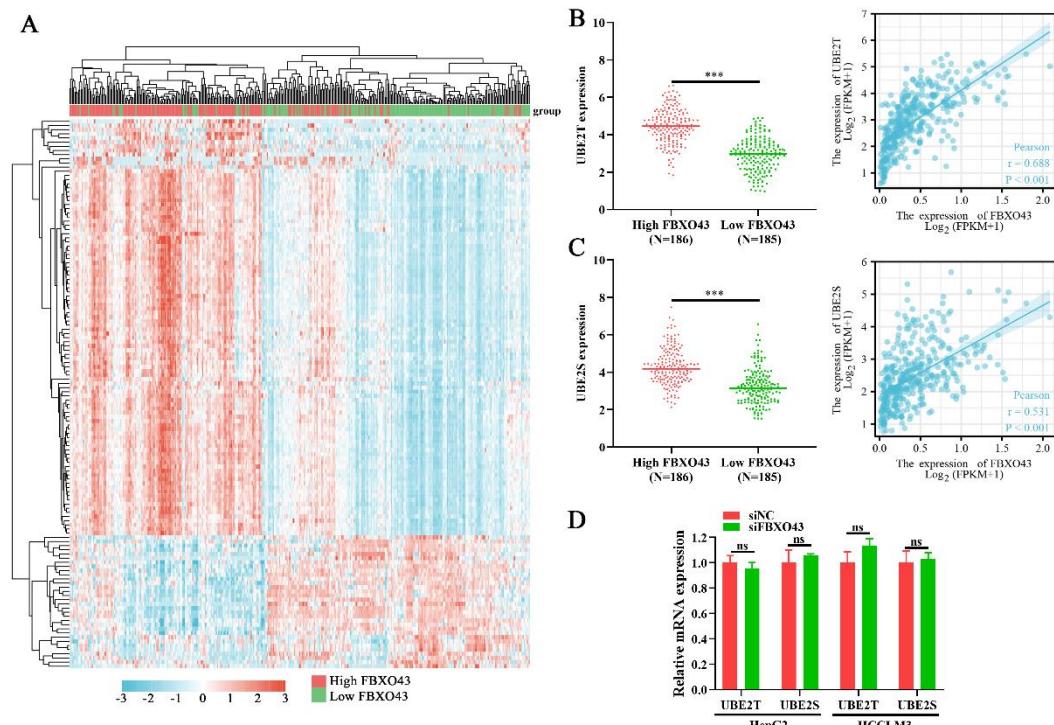


**Supplementary Figure S1. The expression of FBXO43 in subtype of HCC and its prognostic value based on Kaplan-Meier Plotter data.** The expression of FBXO43 in subtype of HCC patients: (A) stage T1 plus T2 vs T3 plus T4; (B) grade G1 plus G2 vs G3 plus G4; (C) AFP ( $\leq 400$  ng/ml) vs AFP ( $>400$  ng/ml). The survival curves from Kaplan-Meier Plotter indicating the prognostic value of FBXO43 including OS (D),

relapsing free survival (E), progression free survival (F), and DSS (G), \*  $p < 0.05$ , \*\*\*,  $p < 0.001$ .

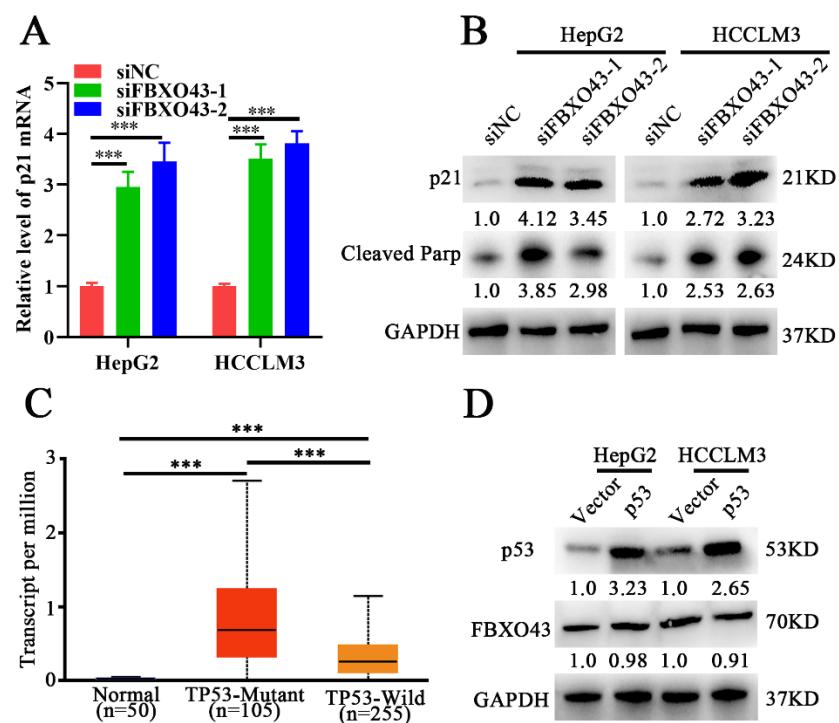


**Supplementary Figure S2. The online analyses of correlation with FBXO43 and prognostic value of M6A writers.** (A) ENCORI data showing IGF2BP2 could bind FBXO43 and negatively regulate its expression in HepG2 cell. (B) and (C) The TCGA data indicating the correlation between IGF2BP2 and M6A writers, including METTL3, METTL14, METTL16, and WTAP, HCC. (D) and (E) The TCGA data indicating the expression and prognostic value of M6A writers in HCC. \*\*\*,  $p < 0.01$ , \*\*\*,  $P < 0.001$ ; ns, no significant difference.

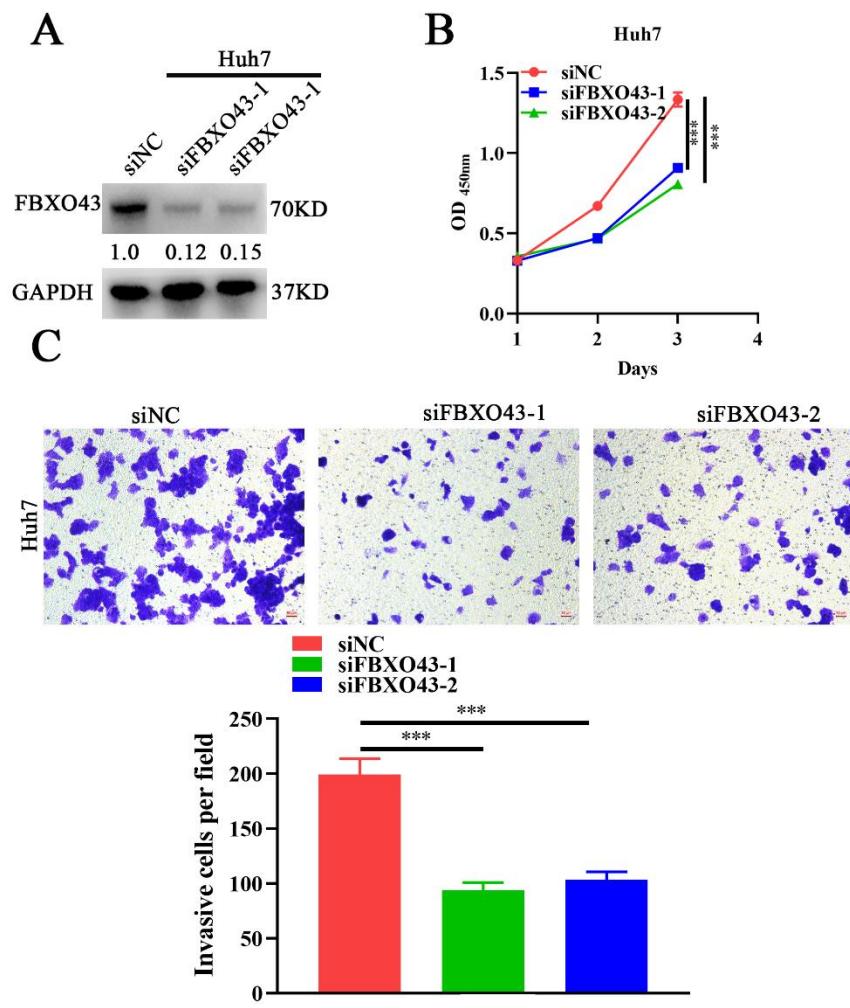


**Supplementary Figure S3. FBXO43 positively correlates with UBE2T and UBE2S, but does not regulate their expression.** (A) Heatmap of differentially expressed genes between High FBXO43 and Low FBXO43 groups. (B) and (C) Scatter plots showing the correlation between UBE2T and UBE2S expression and FBXO43 expression in TCGA data. (D) Bar chart showing relative mRNA expression of UBE2T and UBE2S in HepG2 and HCCLM3 cells under siNC and siFBXO43 conditions. ns, no significant difference.

HCC tissues with low and high FBXO43 expression in TCGA. **(B)** The expression of UBE2T in HCC tissues with low and high FBXO43 and its correlation with FBXO43 in HCC tissues. **(C)** The expression of UBE2S in HCC tissues with low and high FBXO43 and its correlation with FBXO43 in HCC tissues. **(D)** RT-qPCR indicating the expression of UBE2T and UBE2S in HepG2 and HCCLM3 cells with FBXO43 depletion. \*\*\*,  $P < 0.001$ ; ns, no significant difference.



**Supplementary Figure S4. Analyzing the relation between FBXO43 and p53 status in HCC.** **(A)** RT-qPCR indicating the expression of p21 in HepG2 and HCCLM3 cells with FBXO43 depletion. **(B)** WB results indicating the level of p21 and cleaved-Parp in HepG2 and HCCLM3 cells with FBXO43 depletion. **(C)** The UALCAN data showing the expression of TP53 among normal, TP53-mutant, and TP53- wild HCC tissues. **(D)** WB results indicating the level of p53 and FBXO43 in HepG2 and HCCLM3 cells with p53 overexpression. \*\*\*,  $P < 0.001$ .



**Supplementary Figure S5. FBXO43 knockdown inhibits cell growth and invasion in Huh7.** (A)WB results indicating the protein level of FBXO43 in Huh7 cells transfected with siFBXO43s. qRT-PCR. CCK-8 (B) and Transwell (C) assay showing the effects of FBXO43 depletion on the growth and invasion of Huh7 cells. Magnification 100×, Scale bar 50μm. \*\*\*, P < 0.001.

**Supplementary Table S1.** The sequences of primers and siRNAs

Names	Sequences(5'-3')
<b>FBXO43</b>	
Forward	GGAAAGTAAGCAGAAATTGGCGTG
Reverse	GAGTGGCAGCATCCTCGACATT
<b>FBXO43 hnRNA</b>	
Forward	CAGTTGCGAAGAACCAACGTTT
Reverse	AGTTGCCTCCAACGTCTTT
<b>IGF2BP2</b>	
Forward	AGTCCAATTGCATGGGAAAATCA
Reverse	CAACGGCGGTTCTGTGTC
<b>METTL3</b>	
Forward	TTGTCTCCAACCTTCCGTAGT
Reverse	CCAGATCAGAGAGGTGGTAG
<b>UBE2C</b>	
Forward	CTGGCGATAAAGGGATTCTGCC
Reverse	GCGAGAGCTTACCTCAGGTC
<b>UBE2T</b>	
Forward	AGCTGCTCATGTCAGAACCAAAC
Reverse	GTCTGGCATTCTTGAGGAAGGC
<b>UBE2S</b>	
Forward	CCGACACGTACTGCTGACC
Reverse	GCCGCATACTCCTCGTAGTTC
<b>p53</b>	
Forward	CCTCAGCATCTTATCCGAGTGG
Reverse	TGGATGGTGGTACAGTCAGAGC
<b>p21</b>	
Forward	TGTCCGTCAGAACCCATGC
Reverse	AAAGTCGAAGTCCATCGCTC
<b>GAPDH</b>	
Forward	CAGCAAGAGCACAAAGAGGAA
Reverse	ATGGTACATGACAAGGTGCAG
<b>T7-FBOX43 sense</b>	
Forward	TAATACGACTCACTATAGGGATGAGTTAAAGACAAAGAT
Reverse	GAGGCGTTTAAATTCCGCTT
<b>T7-FBOX43 anti-sense</b>	
Forward	TAATACGACTCACTATAGGGAGGCCTTAAATTCCGCTT
Reverse	ATGAGTTAAAGACAAAGATG
<b>siFBOX43-1</b>	CGTGAATTGTTCAAGAT
<b>siFBOX43-2</b>	GCCAAGTTATCAACTTAGAAA
<b>siIGF2BP2</b>	AGTGAAGCTGGAAGCGCATAT
<b>siMETTL3</b>	GCCAAGGAACAATCCATTGTT

**Supplementary Table S2.** The M6A sites of FBXO43 predicted by SRAMP database

Position	Sequence context	Score (binary)	Score (knn)	Score (spectrum)	Score (combined)	Decision(site confidence)
837	ACTGAAGCAGGAAATGGGGCGGA <u>A</u> CTCTCCTCCAATTGTCAACTCC	0.647	0.731	0.517	0.599	High
1102	AGGAAAAGGATAAAACCCCAGAA <u>A</u> CTTTGTGAAACACCTAAAATCA	0.718	0.634	0.593	0.664	Very high
1185	TCTTCGCTCTCTAAAAGGGG <u>AC</u> TTTGAATCACAAAATAGTTCT	0.763	0.879	0.549	0.683	Very high
1331	CACTTTAAAAACAGAAGAAGT <u>GA</u> CTTCATGCAGTCAAAATTGAG	0.566	0.699	0.582	0.579	Moderate
1376	GCTTAATTTCTCAGCAAAAG <u>AC</u> TTCCACAATTGATGATTCAA	0.619	0.758	0.496	0.577	Moderate
1461	ATTCAGGGCAATAATTAAAG <u>ACT</u> CTATCACACATGACTTAGT	0.661	0.723	0.480	0.592	Moderate
1753	ACCAGGAGGGTTCTTTCAAG <u>AA</u> TACTGCAGAACATAAGGGGA	0.563	0.498	0.603	0.576	Moderate
1775	ACTACTGCAGAACATAAGGG <u>AC</u> TCCCAAAGTTGGGGACACCAT	0.639	0.696	0.613	0.631	High
1791	AAGGGGACTCCCAAAGTTGGGG <u>A</u> CACCATAGAAAGACAAGACAT	0.581	0.418	0.611	0.585	Moderate
1831	GACATCTTGGAAAGGTCGAGA <u>AGA</u> CTGTCCACCCCTCGGGAACAAA	0.578	0.507	0.516	0.550	Low
2134	TACTGCAGTGTATACTTG <u>CAGGAC</u> TGATCGGCAAGAAAATGGGT	0.730	0.797	0.645	0.699	Very high

2164	GCAAGAAAATGGGTATA <u>GAAAAAC</u> TGGACATCTTA <u>CAGAATTAA</u>	0.578	0.644	0.674	0.620	High
2169	AAAATGGGTATA <u>GAAAAACTGGAC</u> ATCTTA <u>CAGAATTAAATAT</u>	0.631	0.765	0.653	0.646	High
2650	ACCAGCCATATA <u>AGAAAAGGGGAC</u> TGTGTAGCCGA <u>ACAGCCTGTG</u>	0.759	0.606	0.315	0.574	Moderate
2814	AATT <u>AAAACGCCTCTGAAGAGAC</u> TAA <u>ATAGAACTCCCCATG</u>	0.659	0.547	0.658	0.653	Very high
2826	CTCTGAAGAGACTAA <u>ATAGAAC</u> T <u>CCCCCATGCAGTGTTCTATT</u>	0.561	0.536	0.669	0.603	High
3173	CAGATAATATTAA <u>AGATTTGAAC</u> CTAGACAGAT <u>GGCCTTATAG</u>	0.691	0.685	0.567	0.641	High
3216	AGATTGGAAA <u>ATGAGTACTTGAC</u> TGAA <u>ATGAAATATAAATAAAAA</u>	0.624	0.689	0.570	0.606	High

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**Both uncropped exposure and white pictures were offered as follow:**

**Figure 2B**

**FBXO43**



**GAPDH**

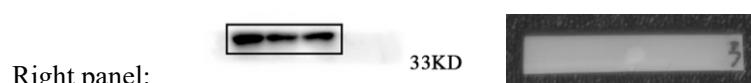


**Figure 2D**

**FBXO43**



**GAPDH**



**Figure 3E**

**IGF2BP2**



**FBXO43**



**GAPDH**





**Figure 3H**

**IGF2BP2**

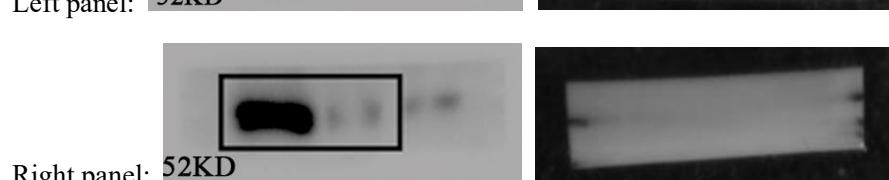
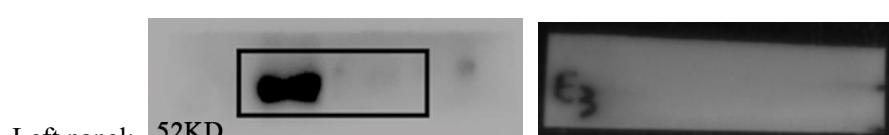


**GAPDH**

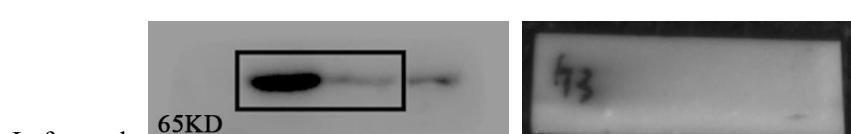


**Figure 3J**

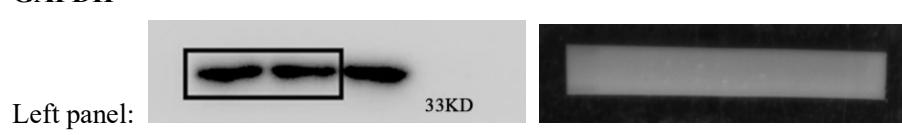
**METTL3**



**FBXO43**



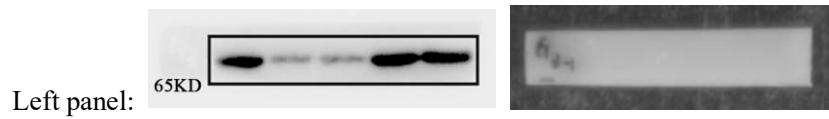
**GAPDH**



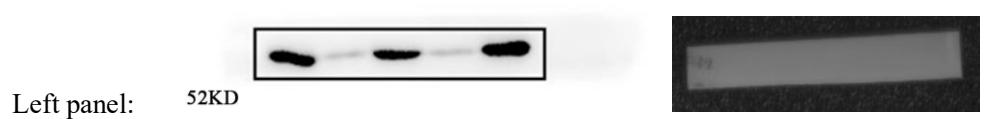


**Figure 4A**

**FBXO43**



**METTL3**



**IGF2BP2**

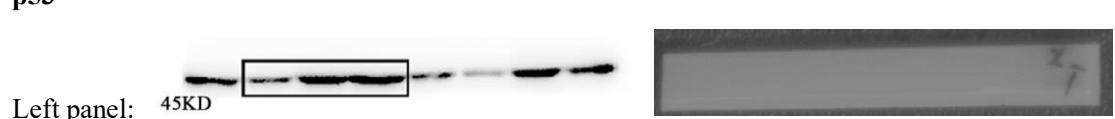


**GAPDH**

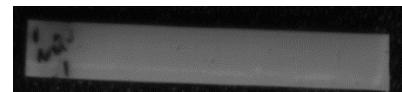
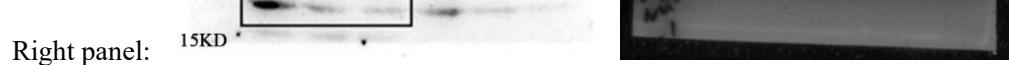
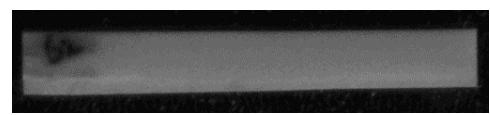
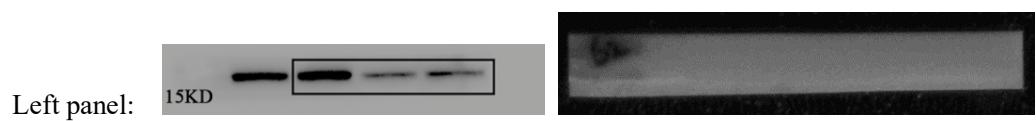


**Figure 5F**

**p53**



**UBE2C**



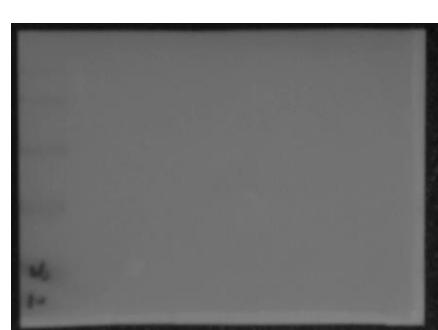
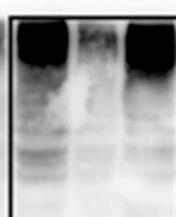
### GAPDH



**Figure 5G**

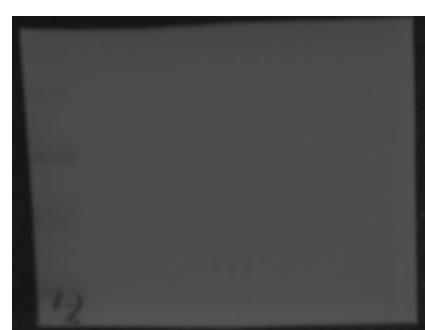
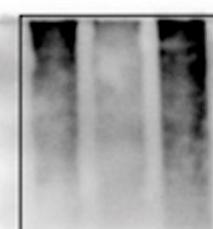
IP

140KD  
53KD  
45KD



Left panel:

140KD  
53KD  
45KD



Right panel:

p53

Left panel:

45KD

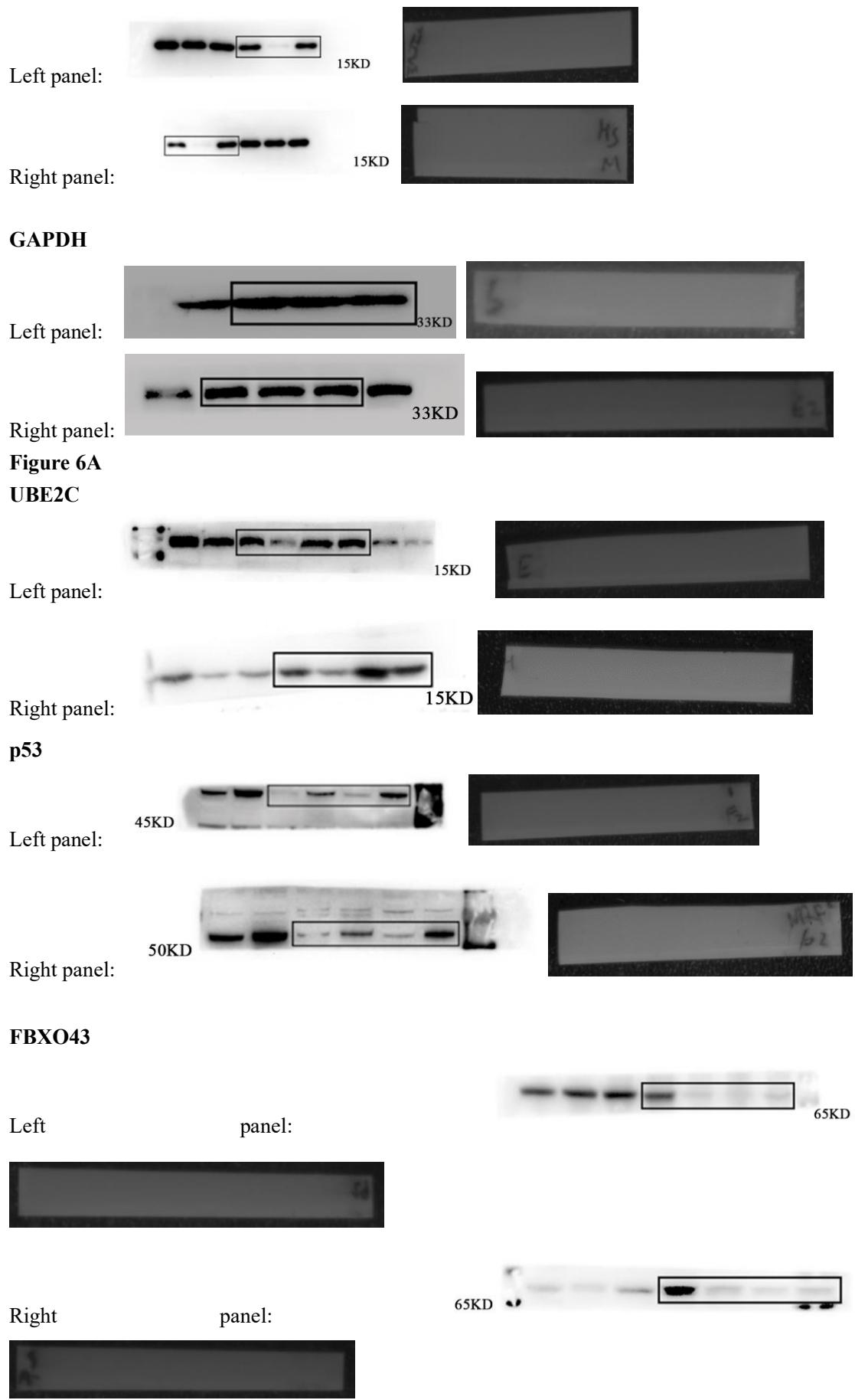


Right panel:

45KD

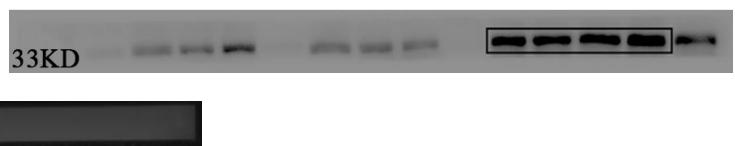


### UBE2C



**GAPDH**

Left panel:



Right panel:

