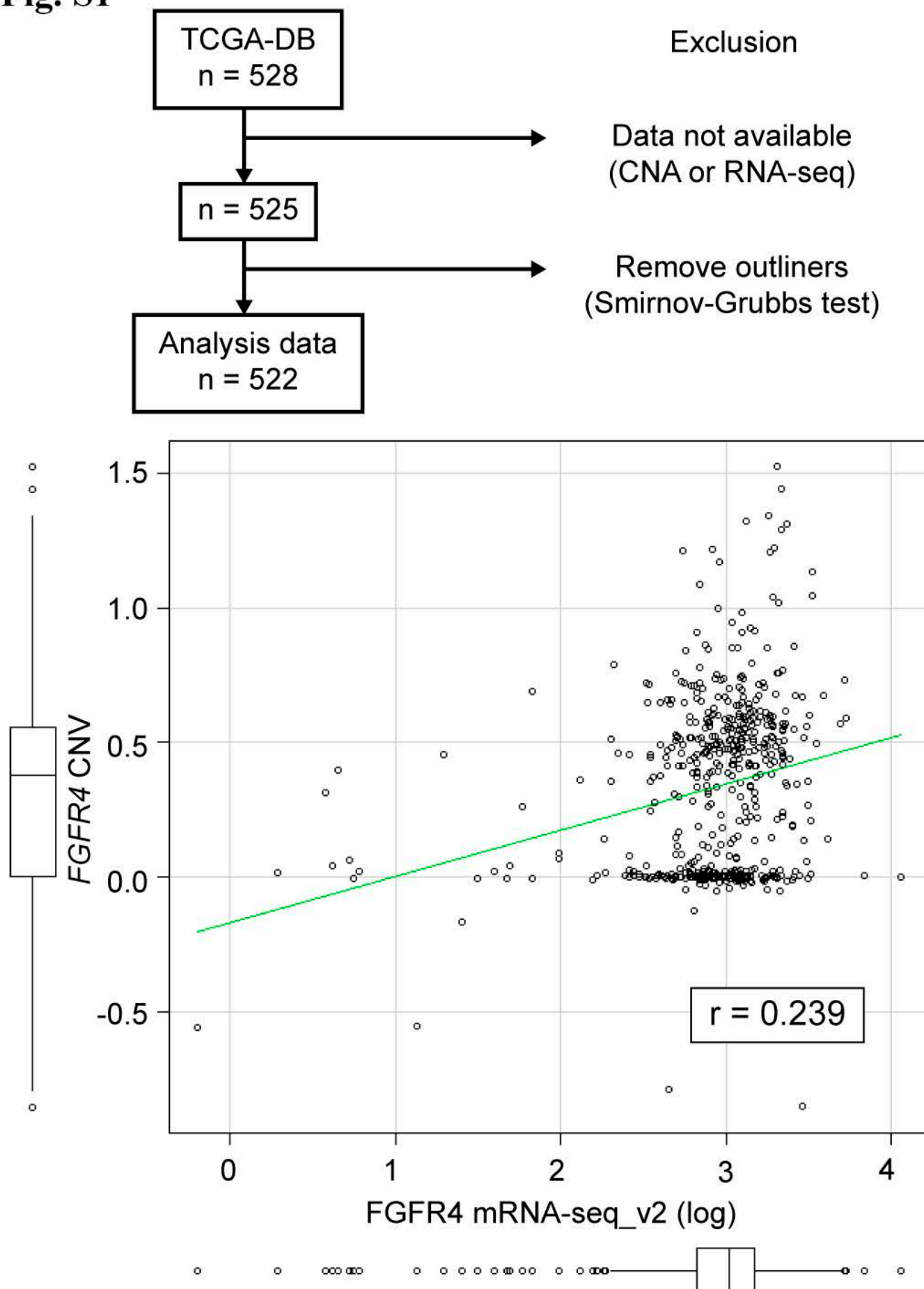


Supplementary Table 1.**Patient characteristics for the analyzed ccRCC specimens**

	Number	%
Total	74	
Median observation time, months (range)	61.15 (0.2–104.2)	
Median age (range)	63.8 (44.2–81.7)	
Sex		
Male	60	81
Female	14	19
Clinical T stage (UICC)		
1a	27	36
1b	16	22
2a	8	11
2b	2	3
3a	15	20
3b	3	4
4	3	4
Clinical N stage (UICC)		
0	66	89
1	5	7
2	3	4
Clinical M stage (UICC)		
0	58	78
1	16	22
Grade (Fuhrman)		
1	15	20
2	25	34
3	26	35
4	8	11
Infiltrative growth (INF)		
α	49	66
β	23	31
γ	1	1
N/A	1	1
Treatment outcome		
Cancer-free survival	48	65
Survival with RCC	8	11
Cancer-specific death	14	19
Death due to other causes	4	5

Fig. S1



Analysis of the relationship between FGFR4 gene copy number and FGFR4 mRNA expression in TCGA database.

Correlation analysis was performed on FGFR4 CNV and FGFR4 mRNA-seq V2 data from 522 cases from TCGA database, excluding outliers for CNV (Smirnov–Grubbs test). Normality was determined using the logarithmic value of RNAseqV2 data.

Pearson's fractional test revealed a weak correlation ($r = 0.239$, $P < 0.01$).

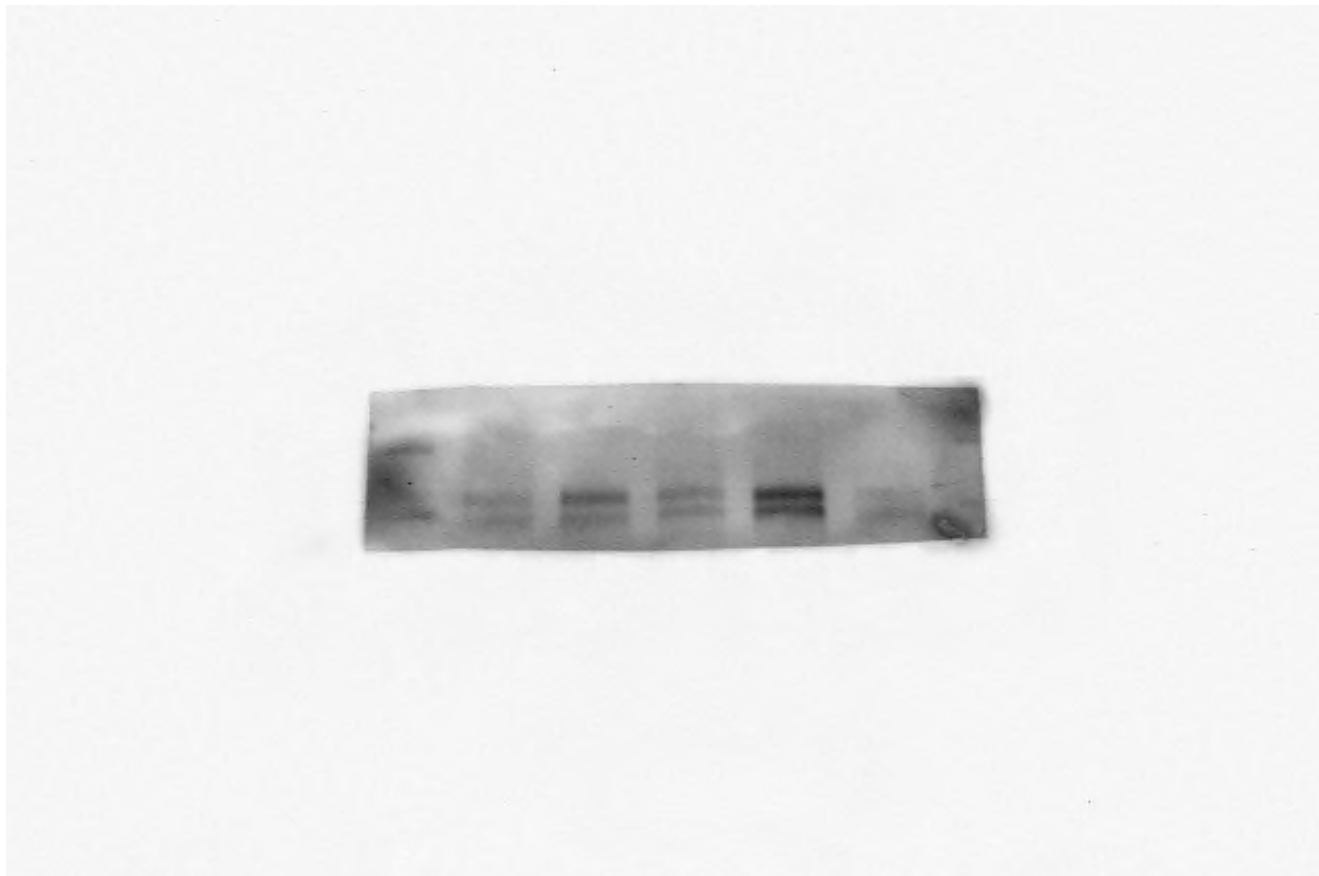


Figure S2 Western blot for FGFR4 protein expression in RCC cell lines.
 From left, molecular weight marker, ACHN, A498, A704, 769-P, 786-O.
 Gel were cut at 70 kDa and other side membrane was used in Western blots for other examination.

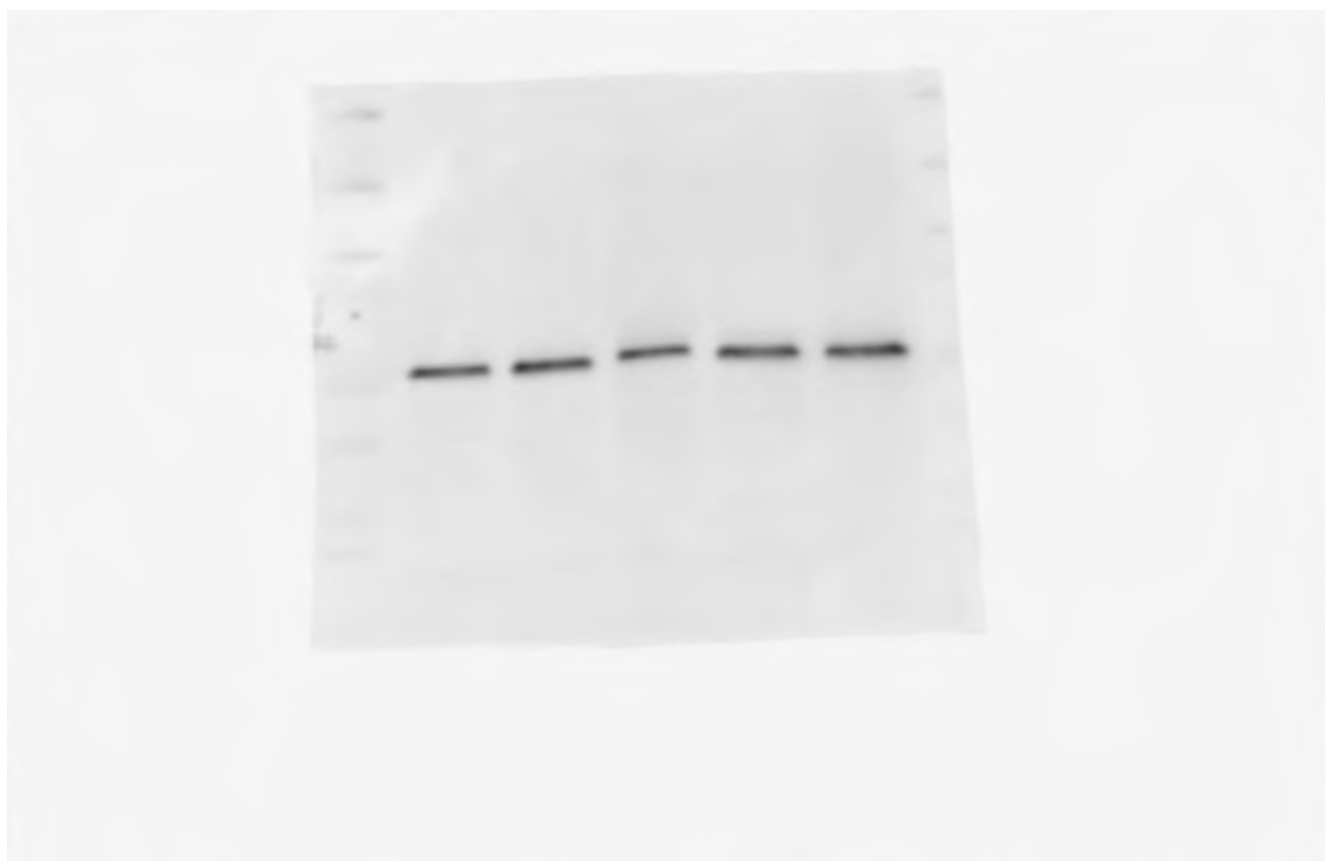


Figure S3 Western blot for α Tubulin protein expression in RCC cell lines. From
 left, molecular weight marker, ACHN, A498, A704, 769-P, 786-O.

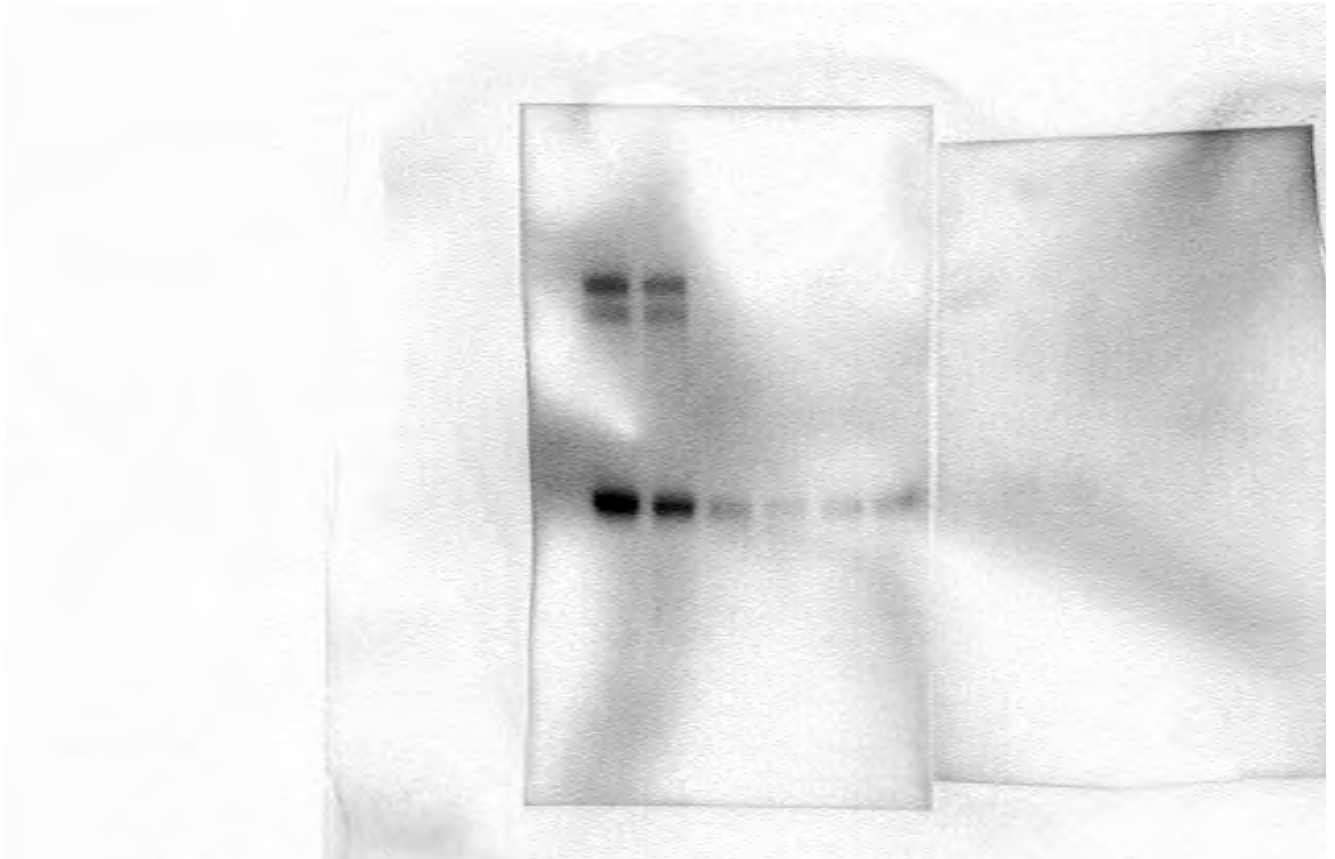


Figure S4 .Western blot for FGFR4 protein expression in A498 cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1(5nM), siFGFR4-2(5nM), siFGFR4-1(25nM), siFGFR4-2(25nM). The right side membrane was external control.

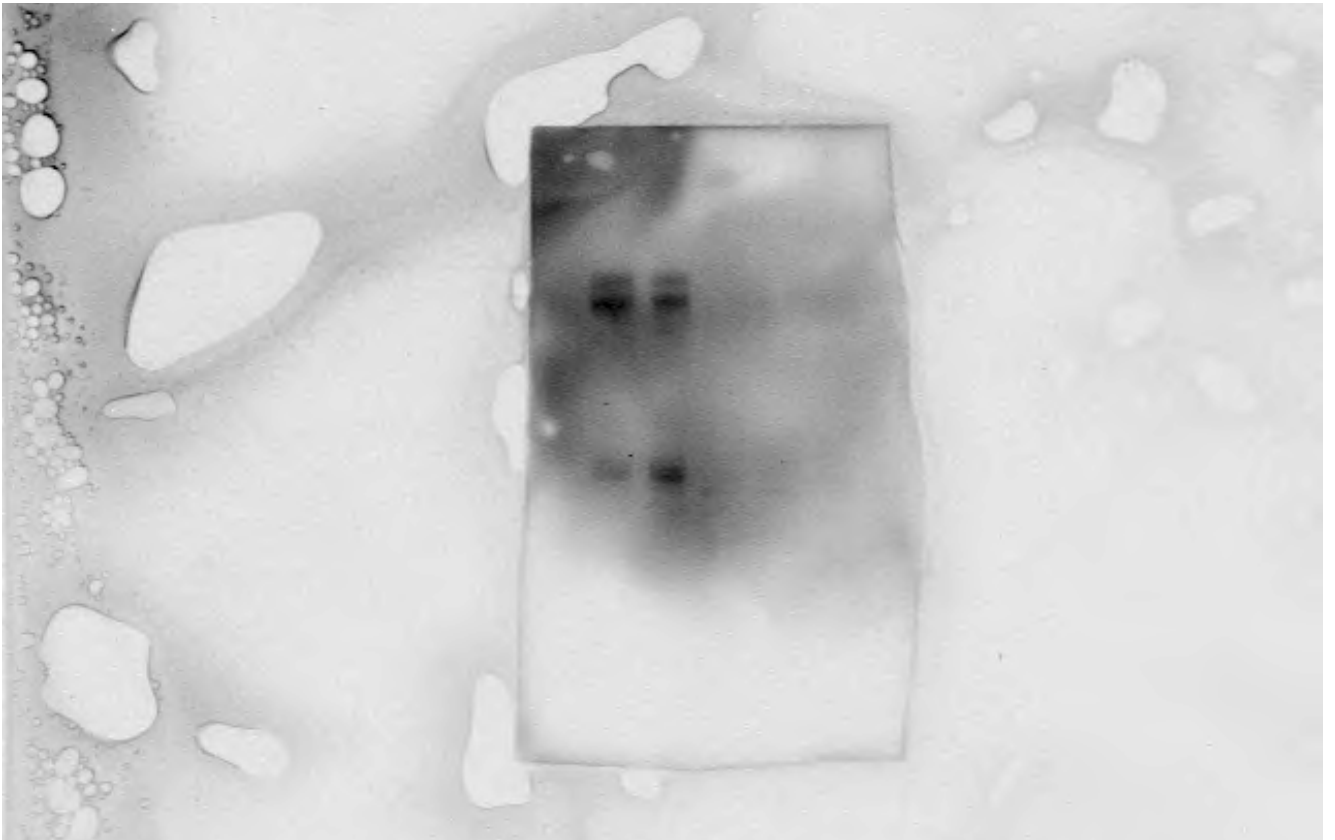


Figure S5 .Western blot for FGFR4 protein expression in A704 cell line after siFGFR4 transfection. From left (right gel) , Control (treatment free), Negative control, siFGFR4-1(5nM), siFGFR4-2(5nM), siFGFR4-1(25nM), siFGFR4-2(25nM).

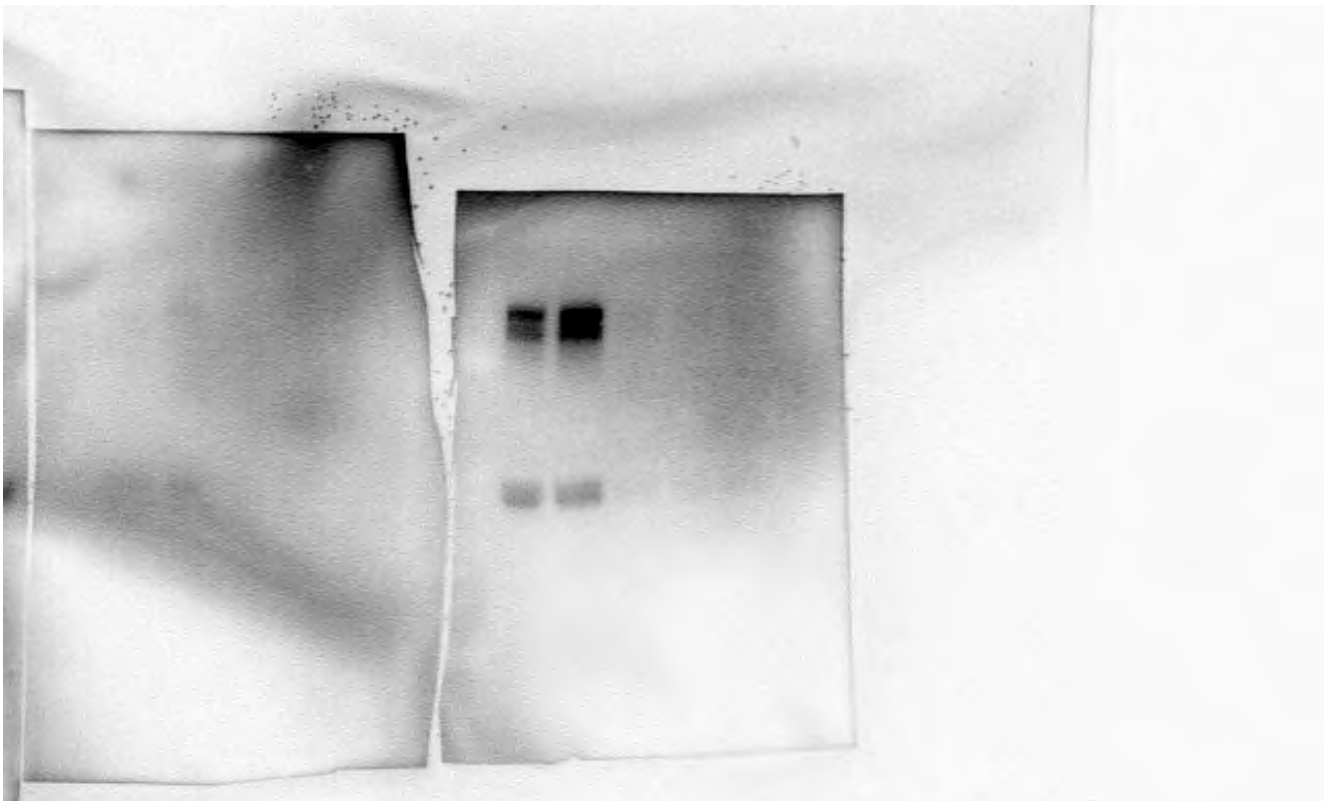


Figure S6

Western blot for FGFR4 protein expression in 769P cell line after siFGFR4 transfection. From left (right membrane) , Control (treatment free), Negative control, siFGFR4-1(5nM), siFGFR4-2(5nM), siFGFR4-1(25nM), siFGFR4-2(25nM). the left side membrane was external control.

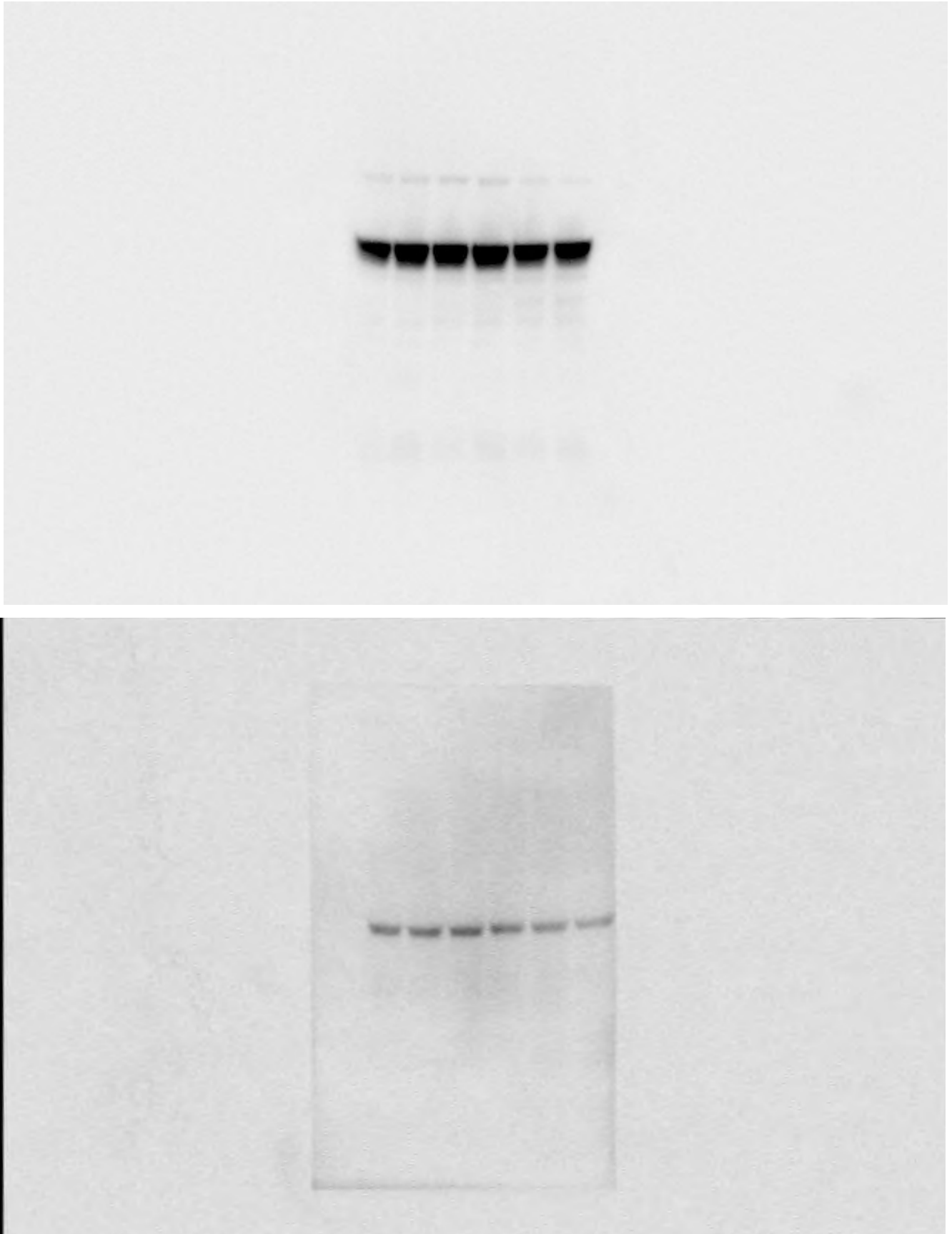


Figure S7 Western blot for AKT(Upper gel) and Phospho-AKT(Ser473, lower gel) protein expression in A498 cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2(5nM), siFGFR4-1 (25nM), siFGFR4-2(25nM).

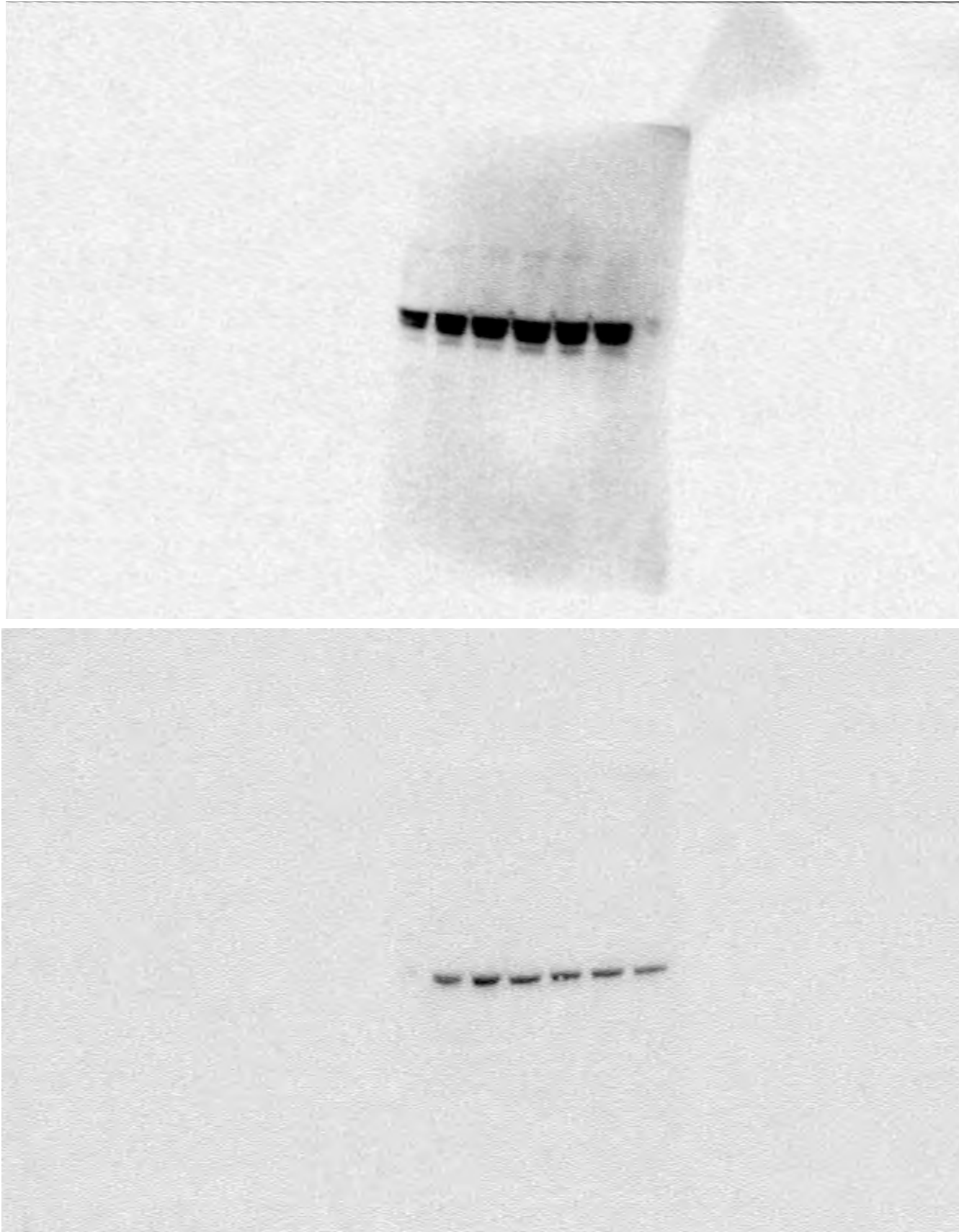


Figure S8 Western blot for AKT(Upper gel) and Phospho-AKT(Ser473, lower gel) protein expression in A704 cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1(5nM), siFGFR4-2(5nM), siFGFR4-1(25nM), siFGFR4-2(25nM).

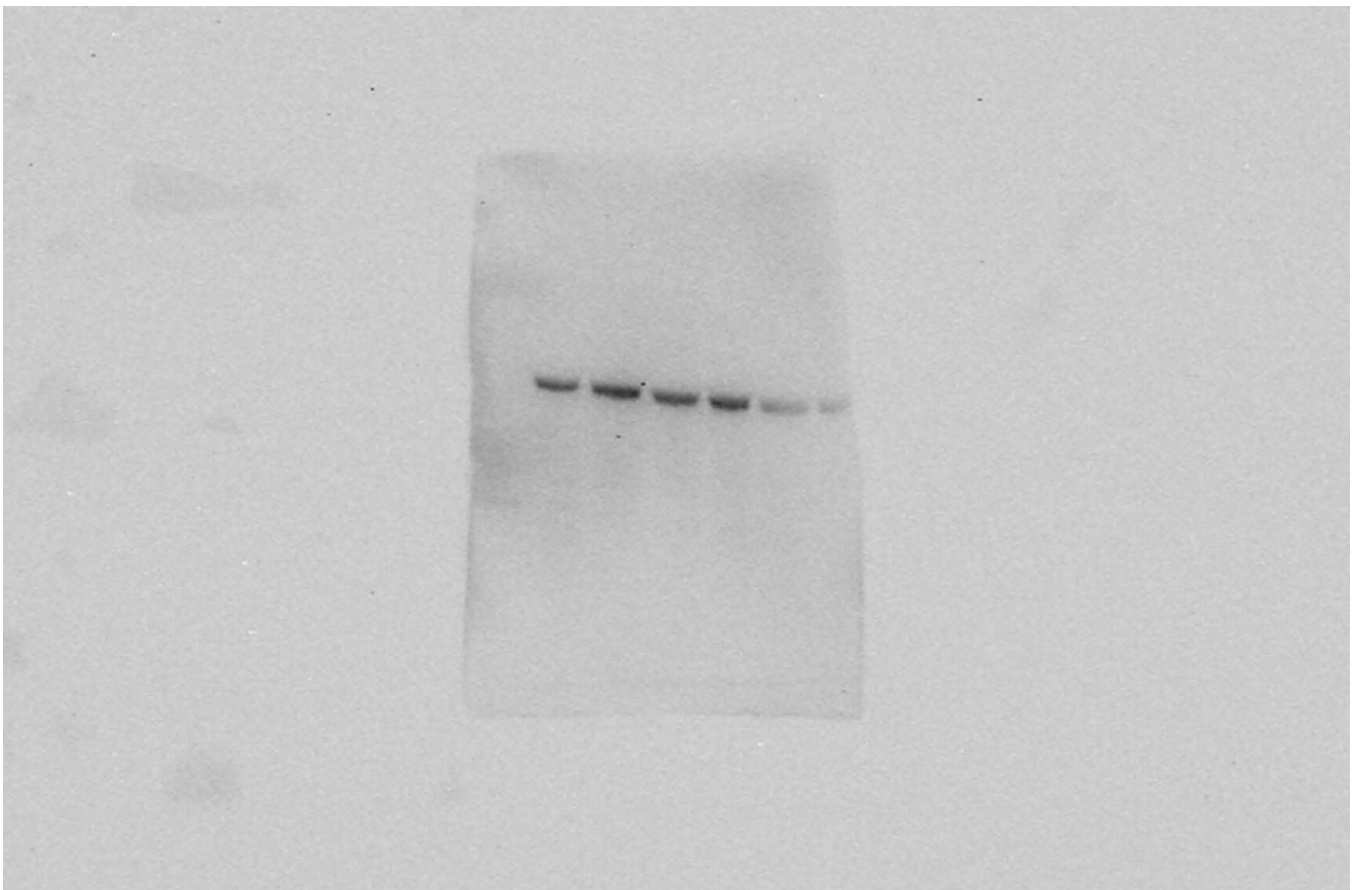
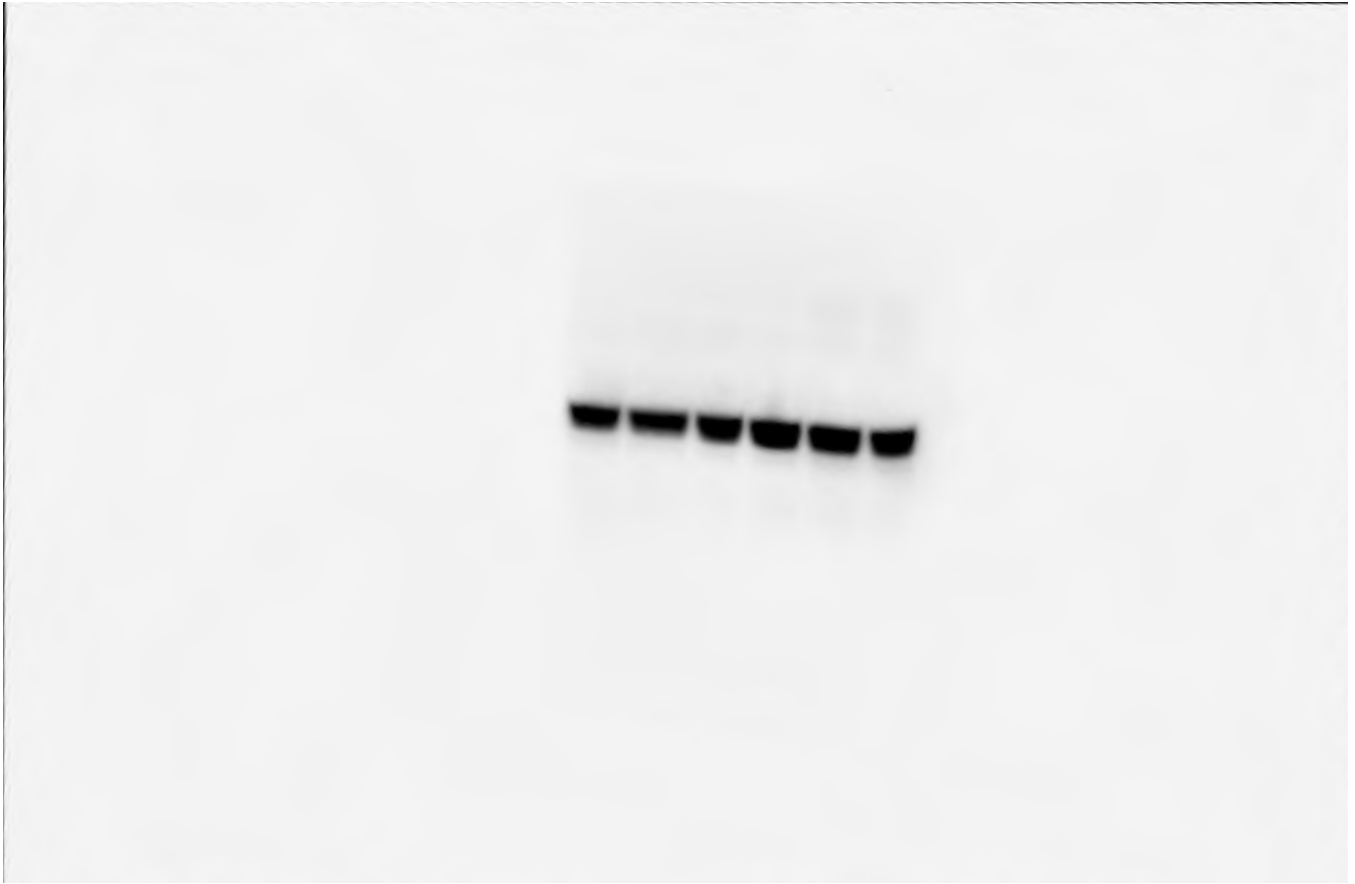


Figure S9 Western blot for AKT(Upper gel) and Phospho-AKT(Ser473, lower gel) protein expression in 769P cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2(5nM), siFGFR4-1 (25nM), siFGFR4-2(25nM).

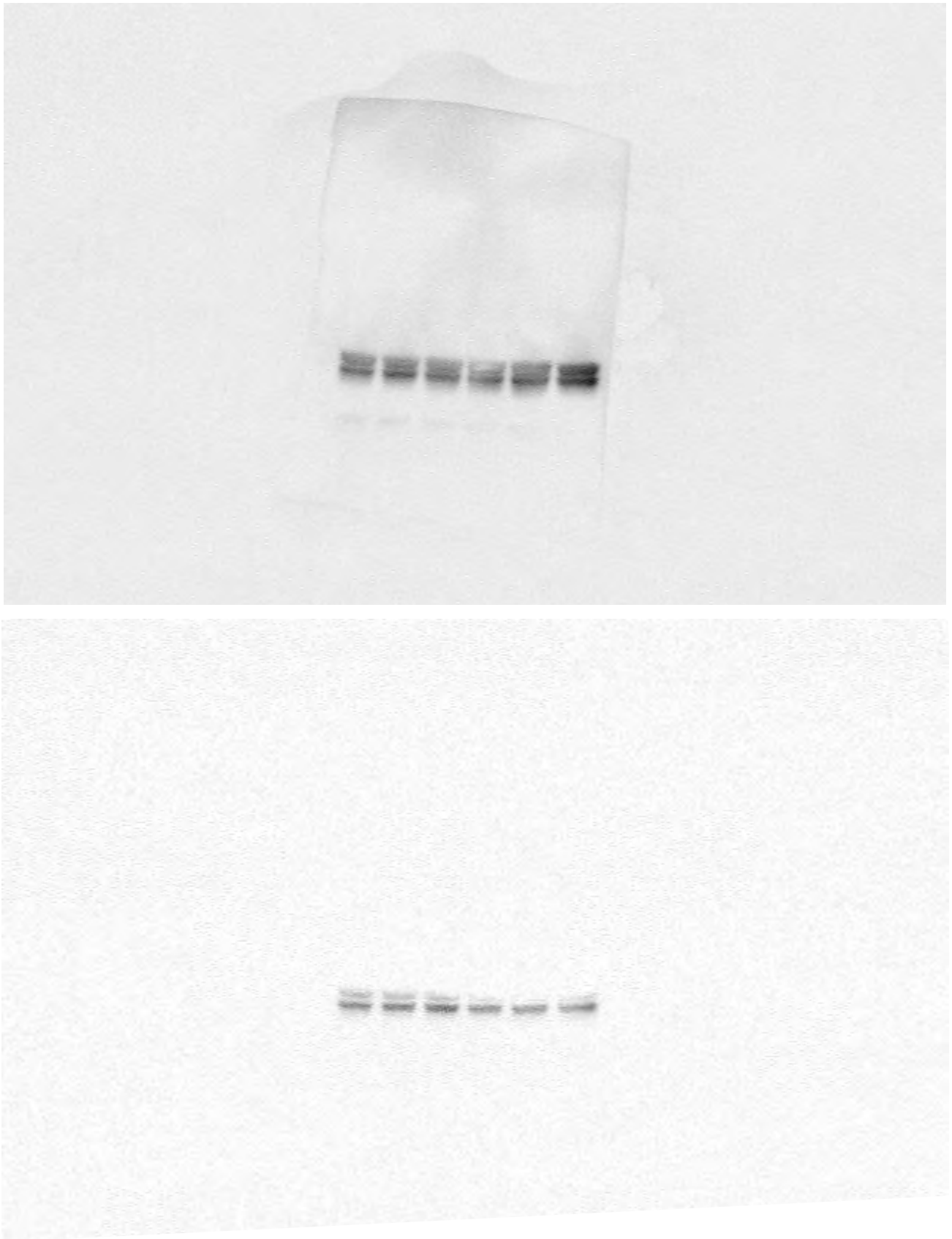


Figure S10 Western blot for ERK1/2 (Upper gel) and Phospho-ERK1/2 (Thr202/Tyr204, lower gel) protein expression in A498 cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2 (5nM), siFGFR4-1 (25nM), siFGFR4-2 (25nM).

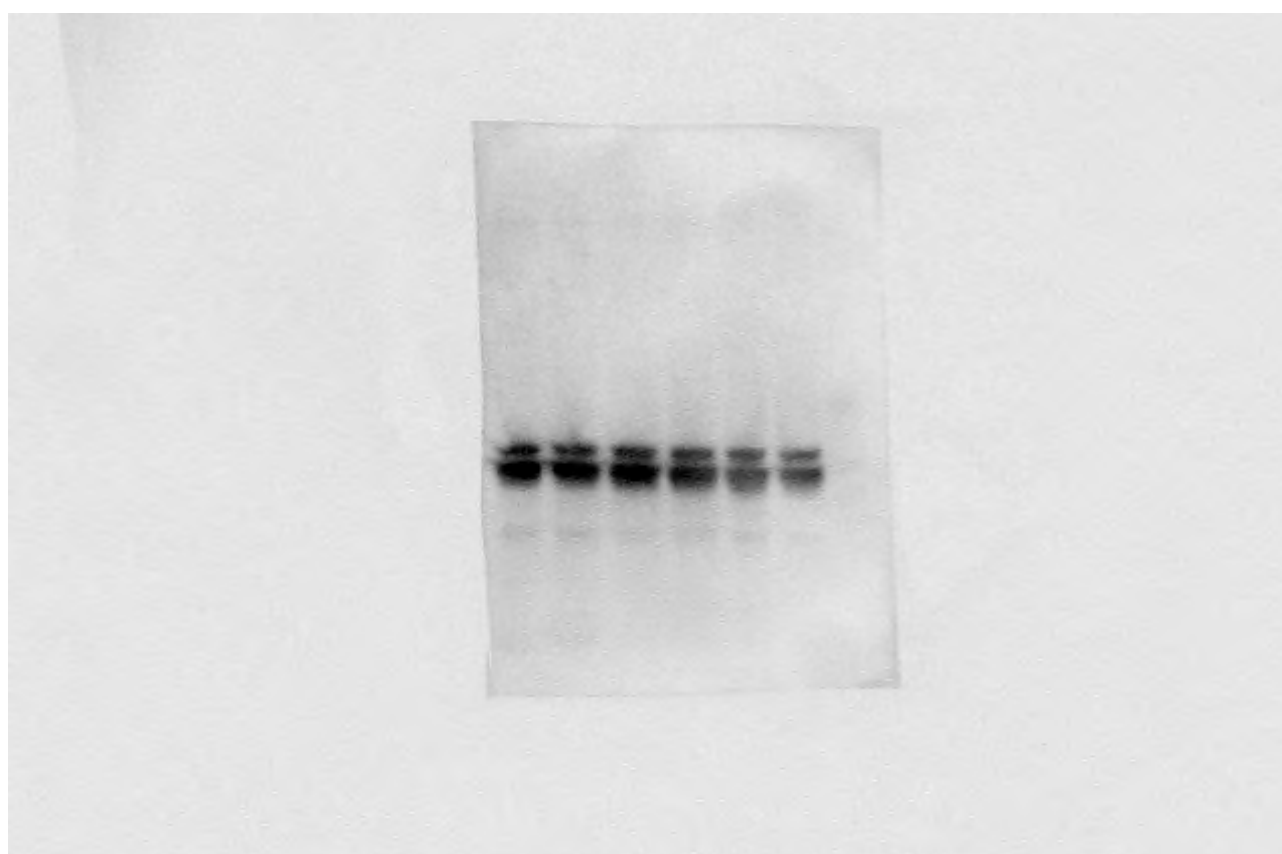
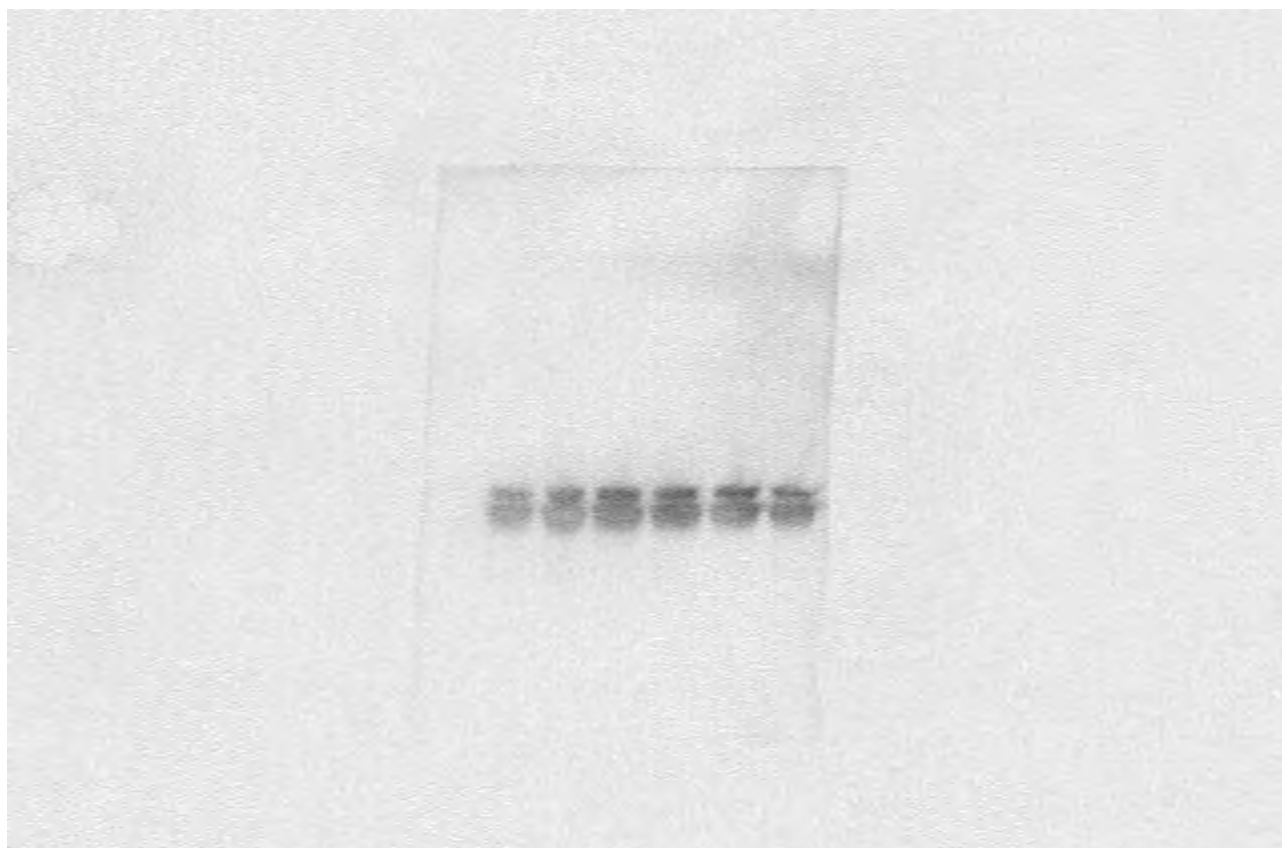


Figure S11 Western blot for ERK1/2 (Upper gel) and Phospho-ERK1/2 (Thr202/Tyr204, lower gel) protein expression in A704 cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2 (5nM), siFGFR4-1 (25nM), siFGFR4-2 (25nM).

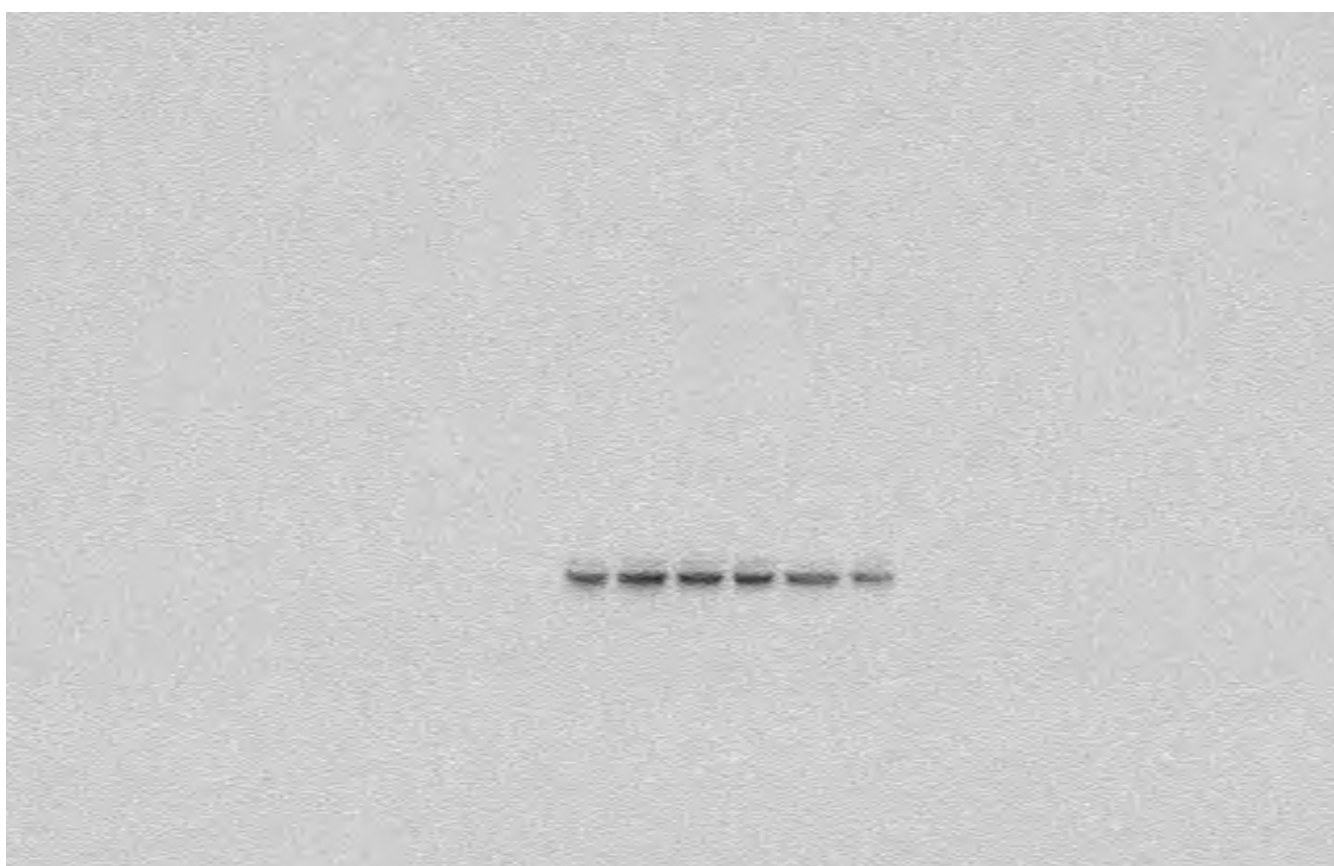
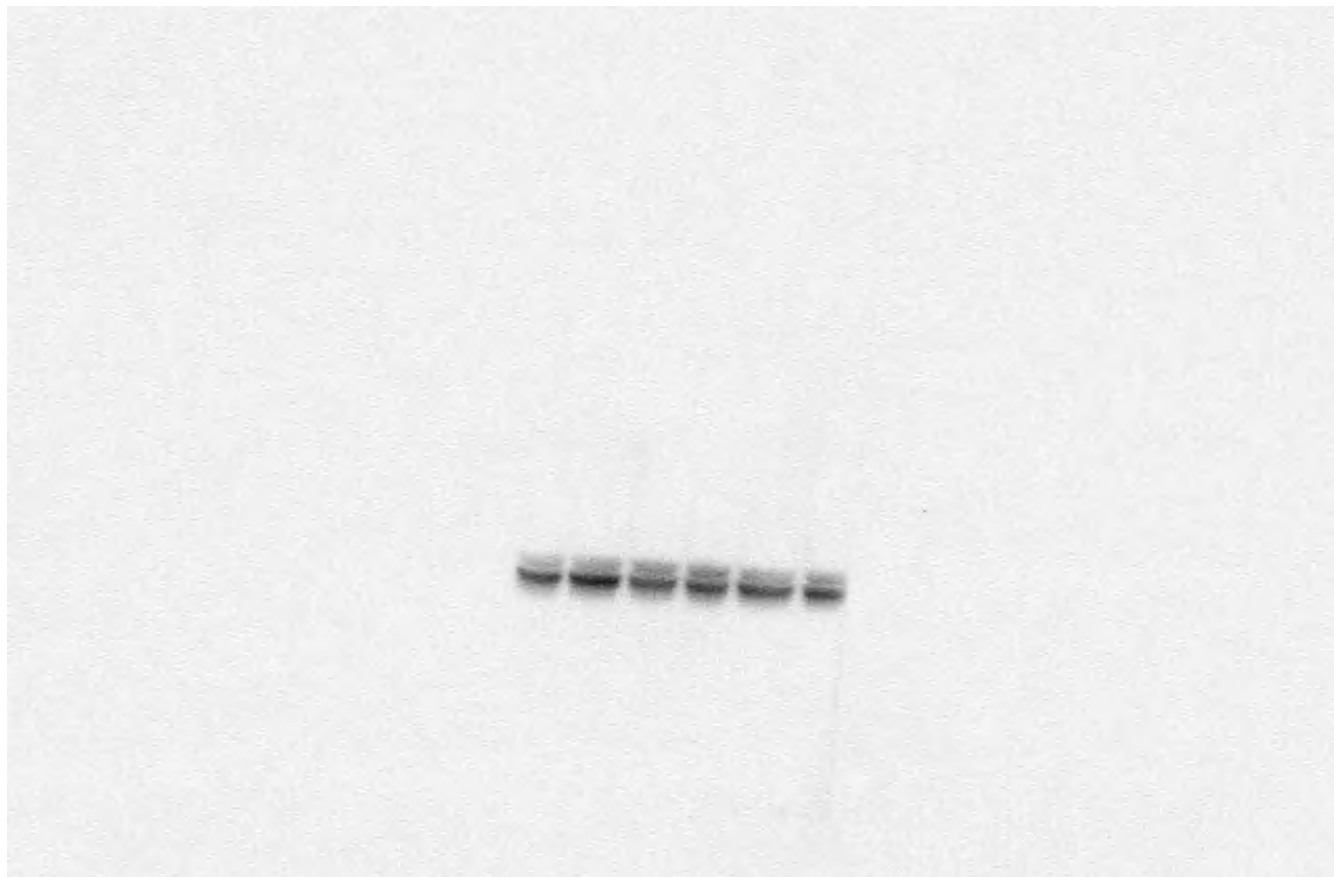


Figure S12

Western blot for ERK1/2 (Upper gel) and Phospho-ERK1/2 (Thr202/Tyr204, lower gel) protein expression in 769P cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2(5nM), siFGFR4-1 (25nM), siFGFR4-2(25nM).

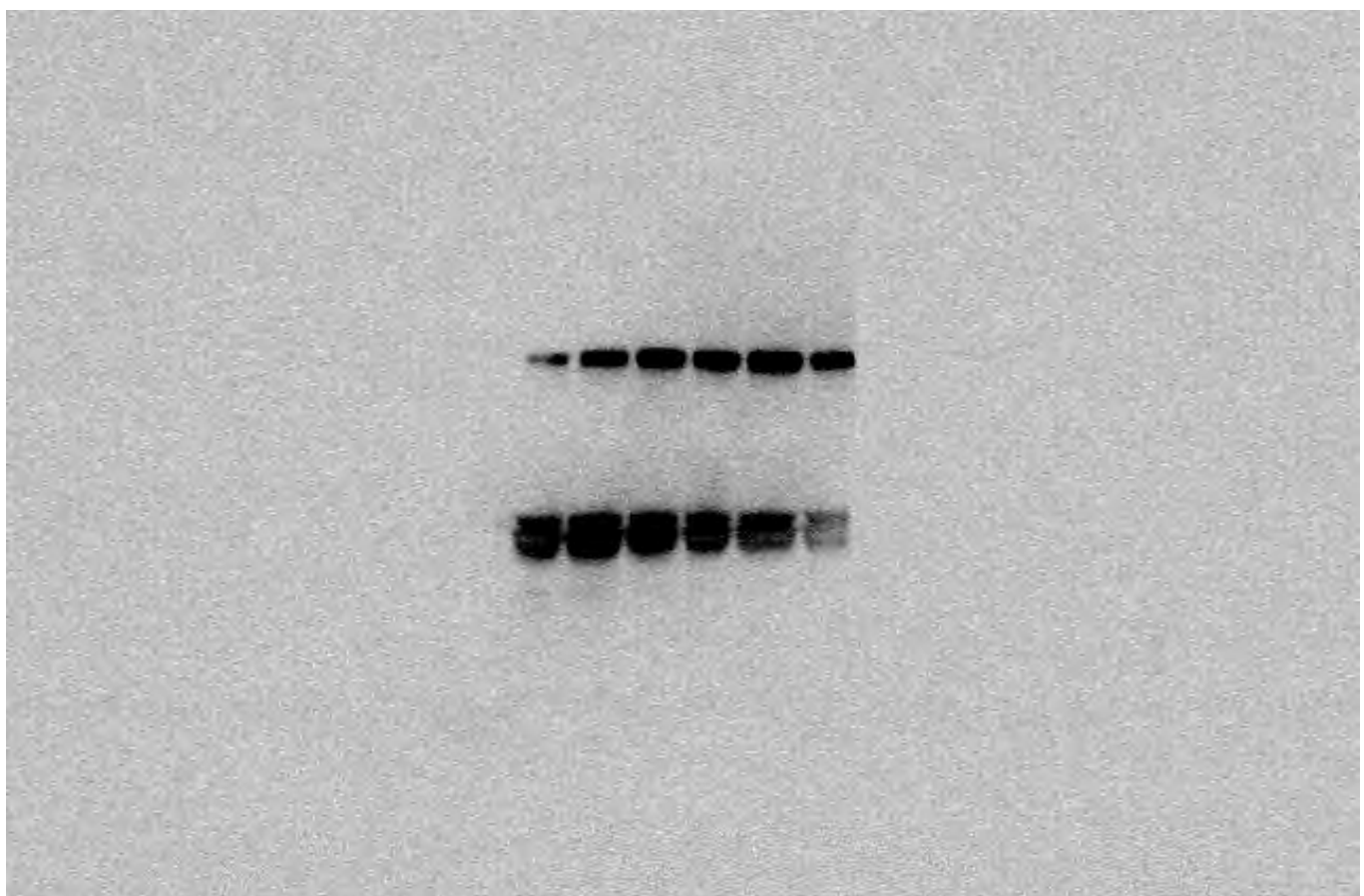


Figure S13 Western blot for STAT3 (Upper gel) and Phospho-STAT3 (Tyr705, lower gel) protein expression in A498 cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2 (5nM), siFGFR4-1 (25nM), siFGFR4-2 (25nM). The lower band of each gel is a remnant of Phosho-ERK1/2 experiment.

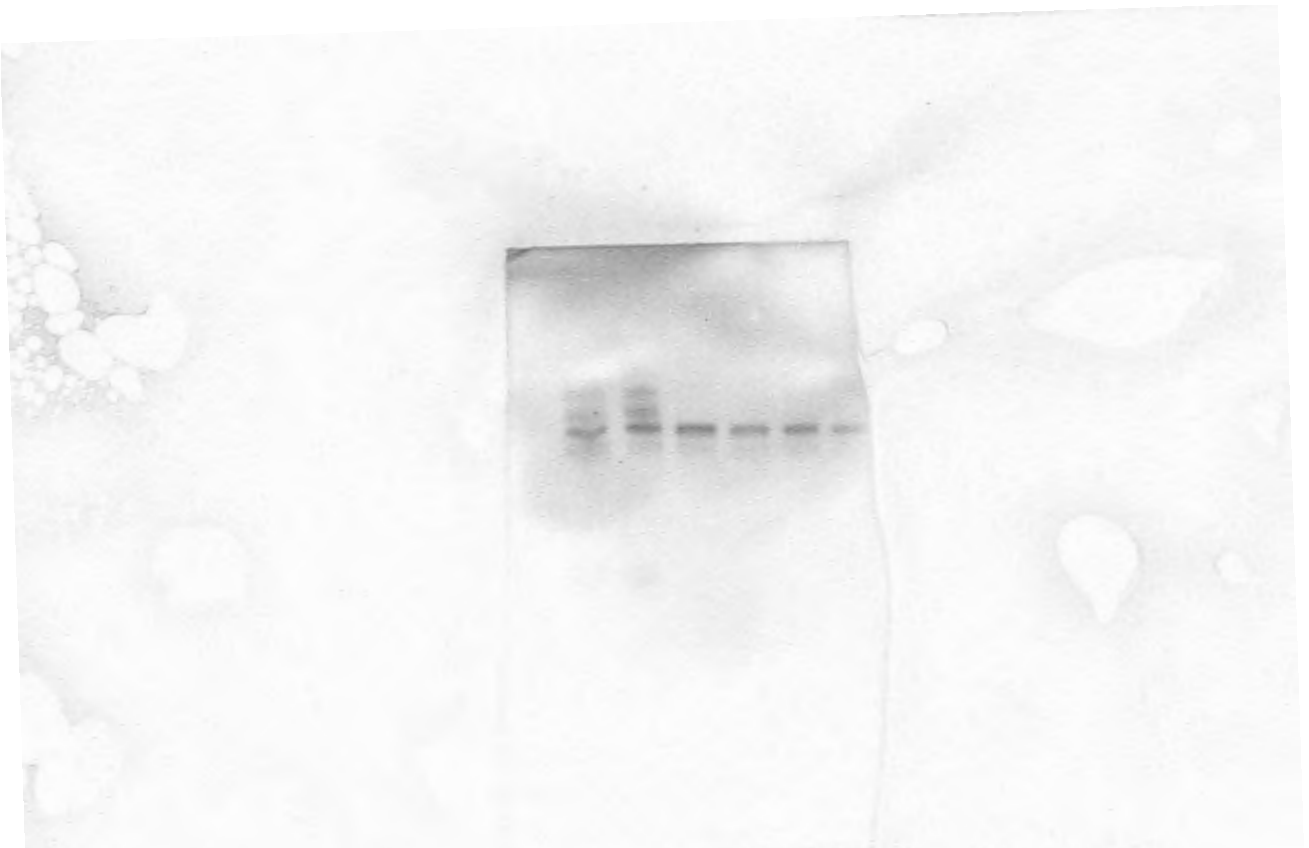
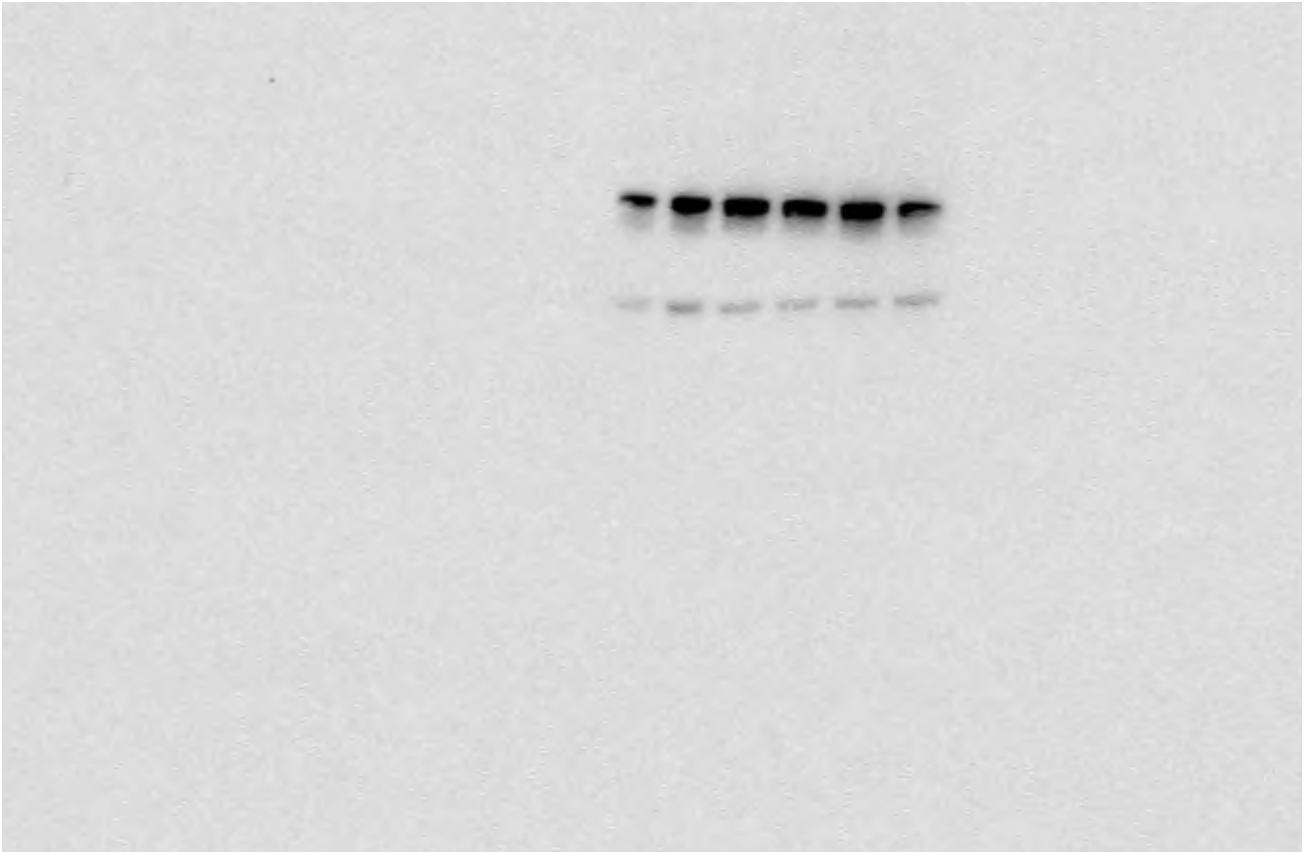


Figure S14 Western blot for STAT3 (Upper gel) and Phospho-STAT3 (Tyr705, lower gel) protein expression in A704 cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2(5nM), siFGFR4-1 (25nM), siFGFR4-2(25nM).

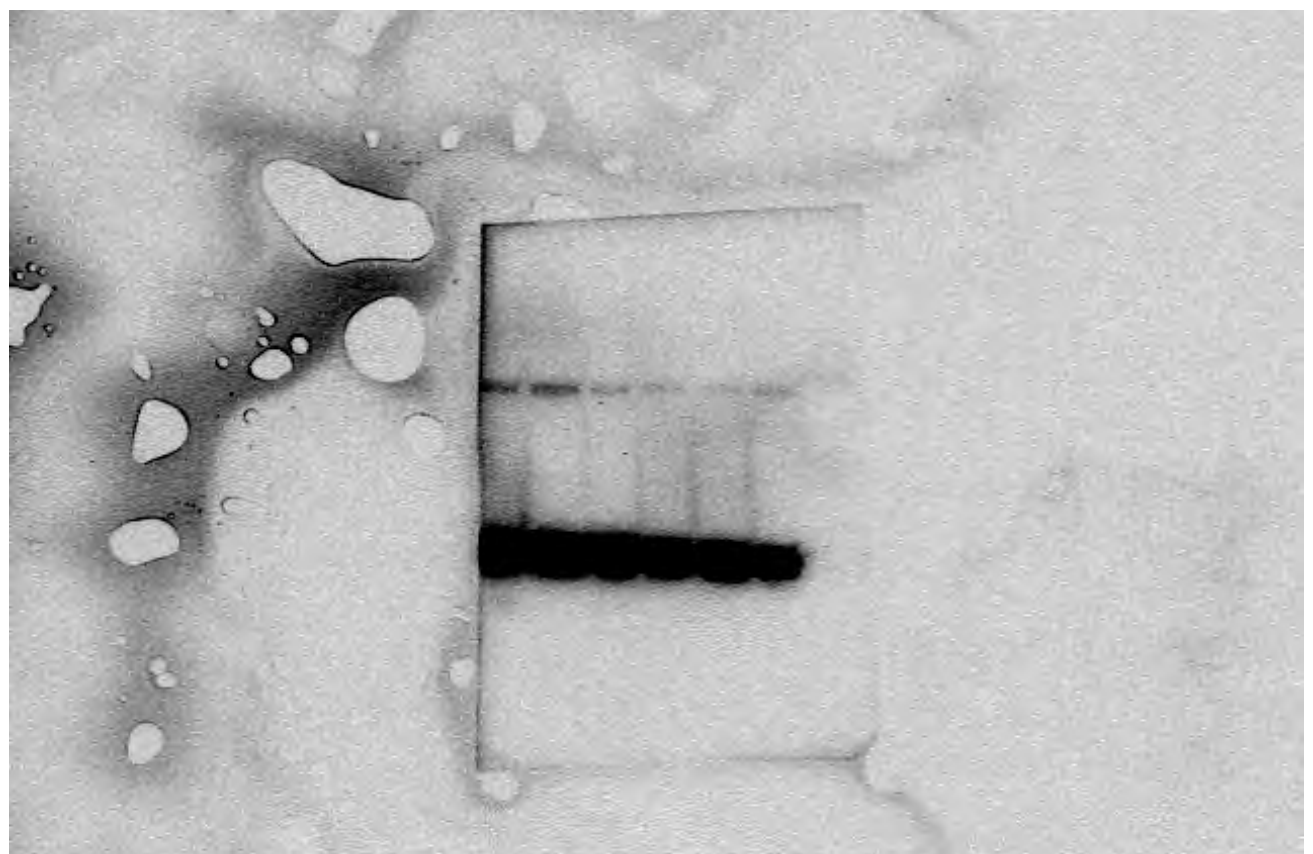
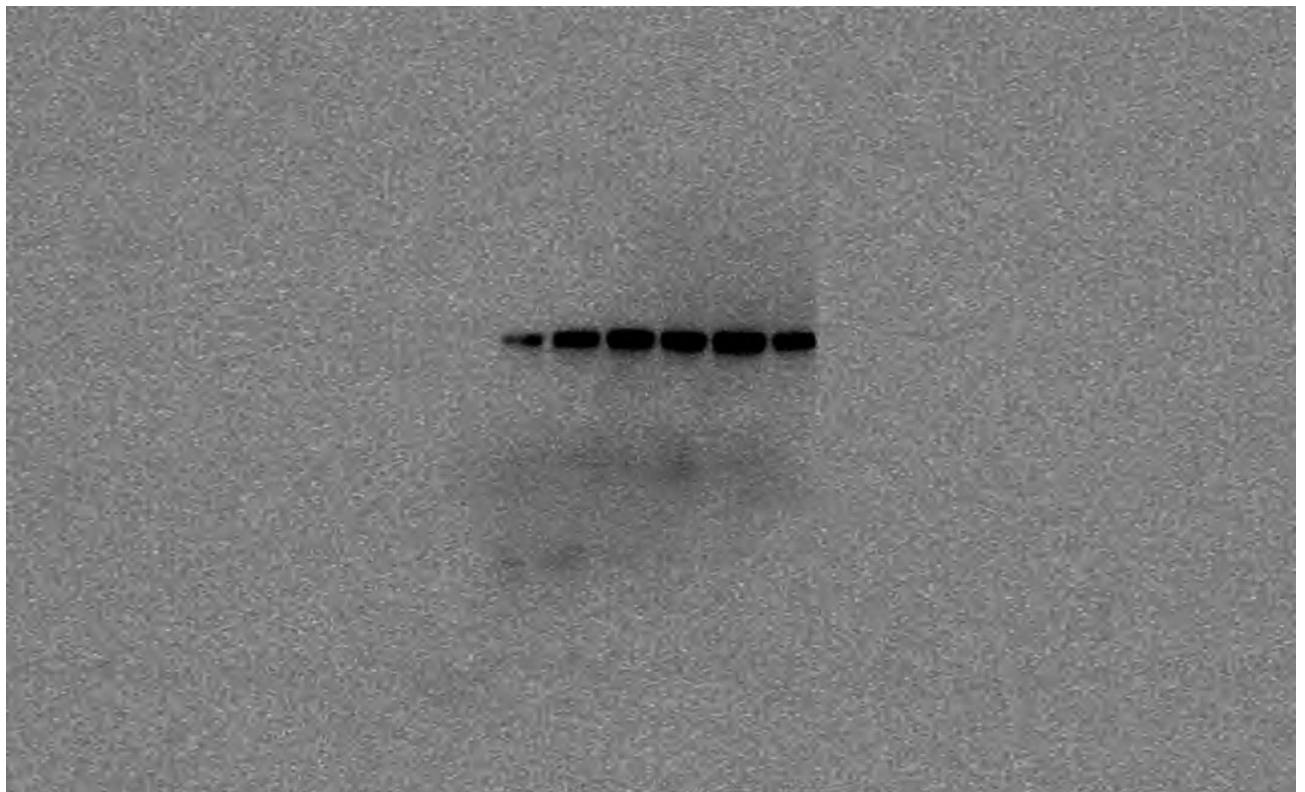


Figure S15 Western blot for STAT3 (Upper gel) and Phospho-STAT3 (Tyr705, lower gel) protein expression in 769P cell line after siFGFR4 transfection. From left, Control (treatment free), Negative control, siFGFR4-1 (5nM), siFGFR4-2 (5nM), siFGFR4-1 (25nM), siFGFR4-2 (25nM).

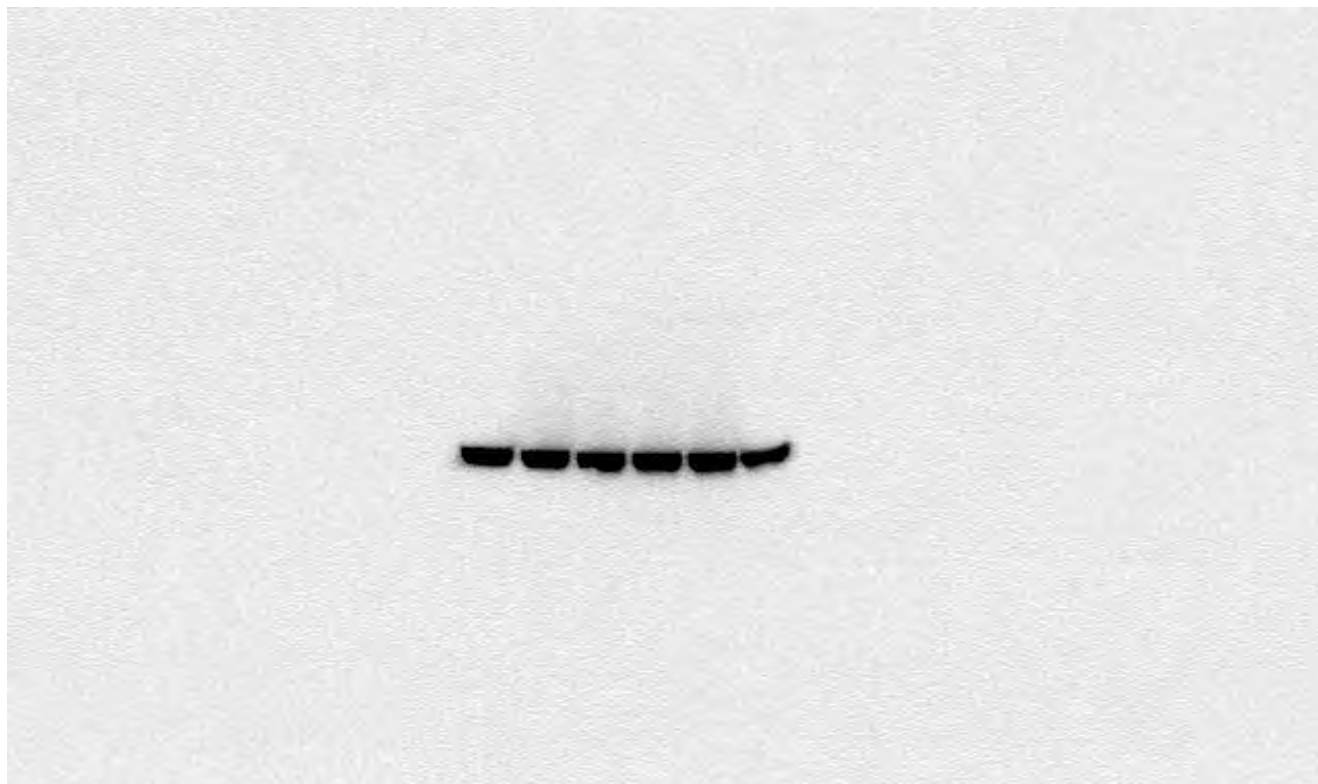


Figure S16 Western blot for α Tubulin expression of A498 cell after siFGFR4 transfection.

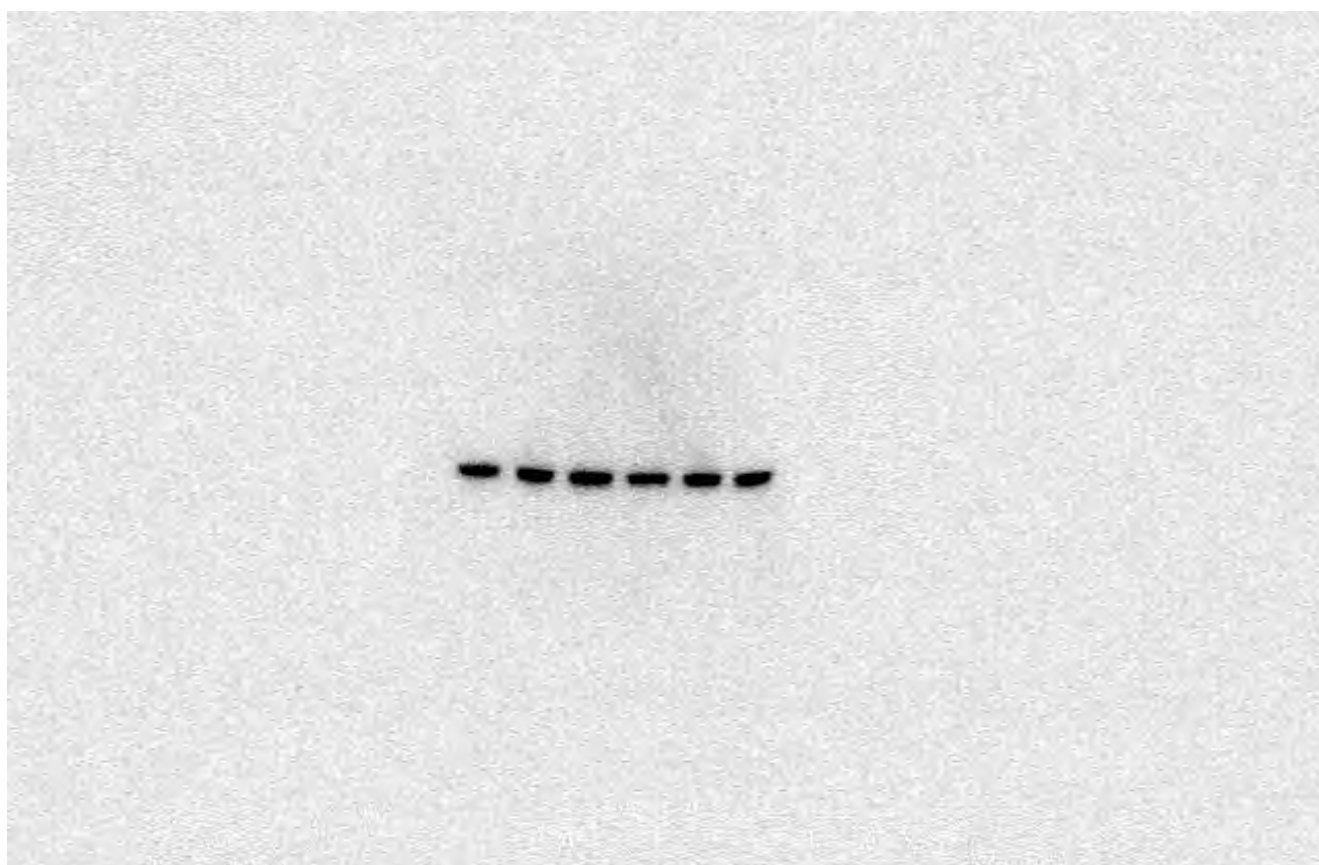


Figure S17 Western blot for α Tubulin expression of A704 cell after siFGFR4 transfection.



Figure S18 Western blot for α Tubulin expression of 769P cell after siFCFR4 transfection.



Figure S19 Western blot for phospho-FGFR(Thr653/654) of A498 cell line after BLU9931 treatment. From left, MW marker, Control, 1 nM, 10 nM, 100 nM, 1 μ M, 10 μ M of BLU9931 treatment concentration. Gel were cut at 70 kDa and other side membrane was used in Western blots for other examination.

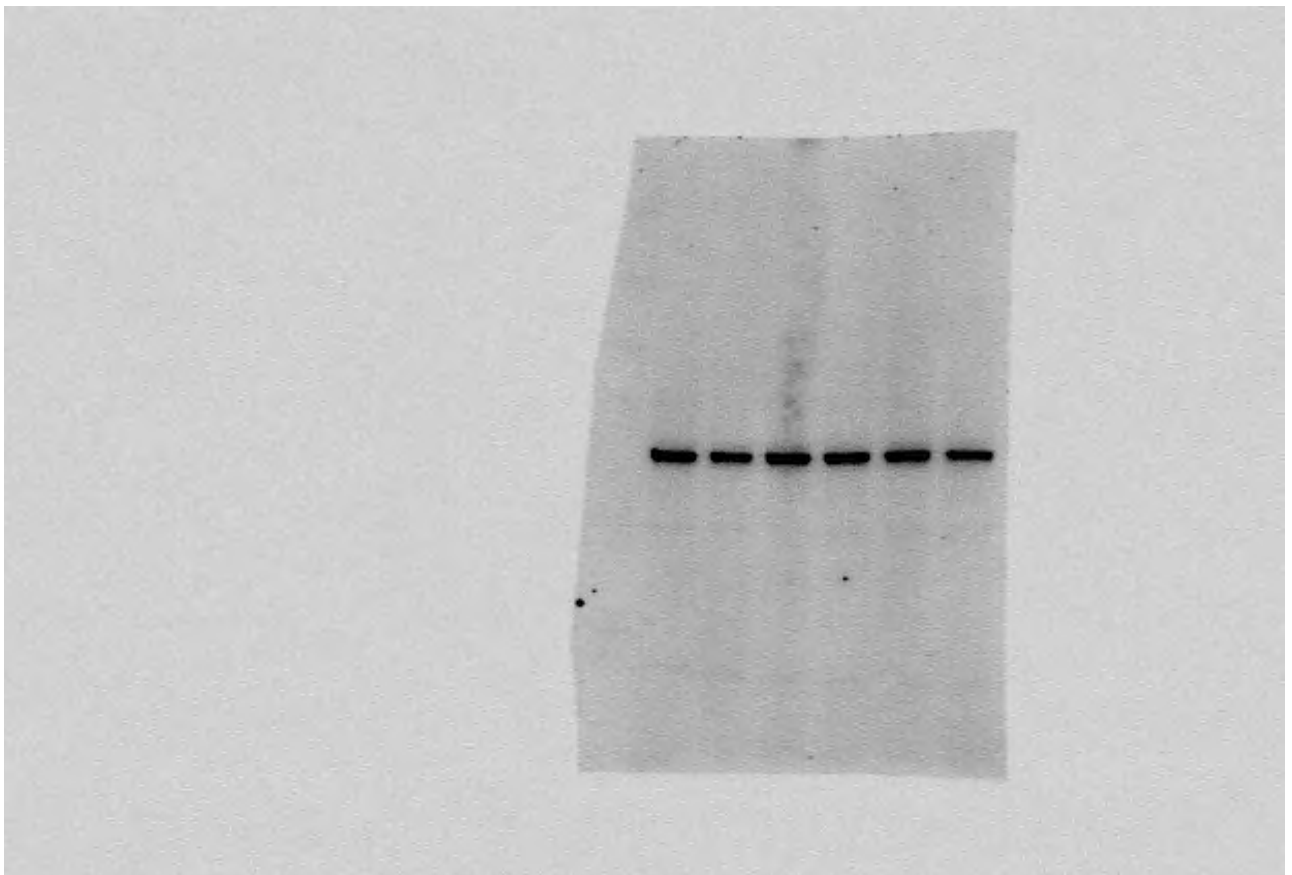


Figure S20 Western blot for α -Tubulin of A498 cell line after BLU9931 treatment. From left, Control, 1 nM, 10 nM, 100 nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

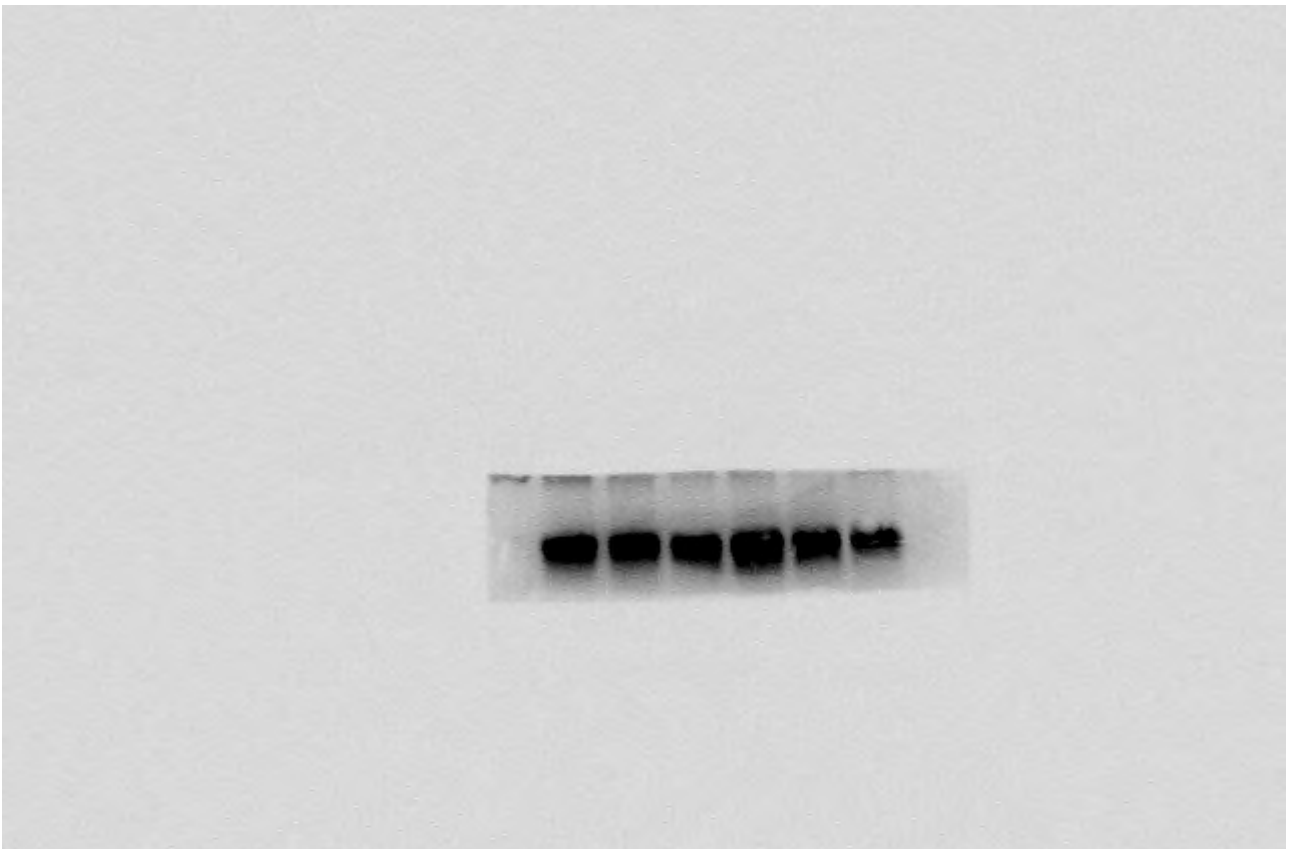


Figure S21 Western blot for AKT of A498 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

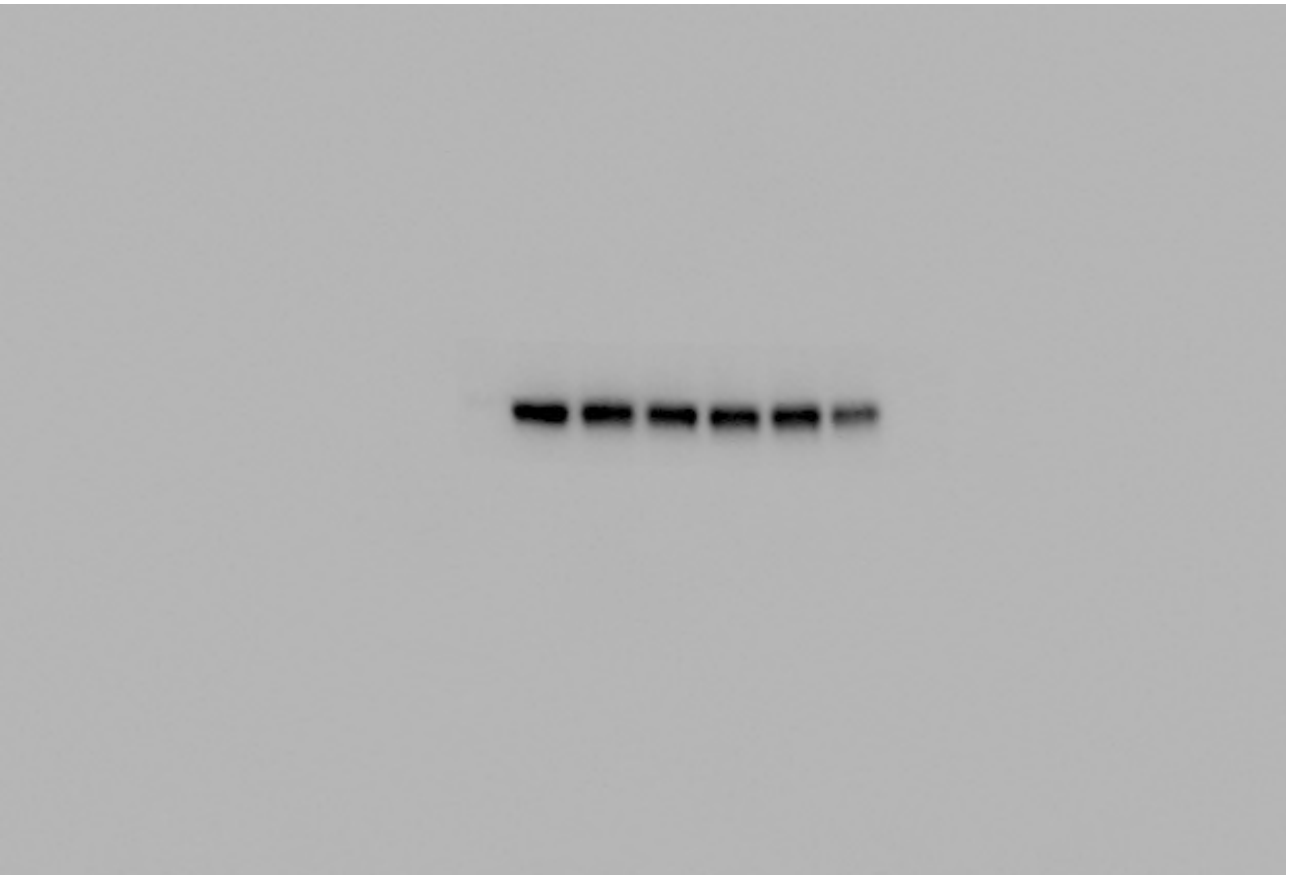


Figure S22 Western blot for phospho-AKT(Ser473) of A498 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

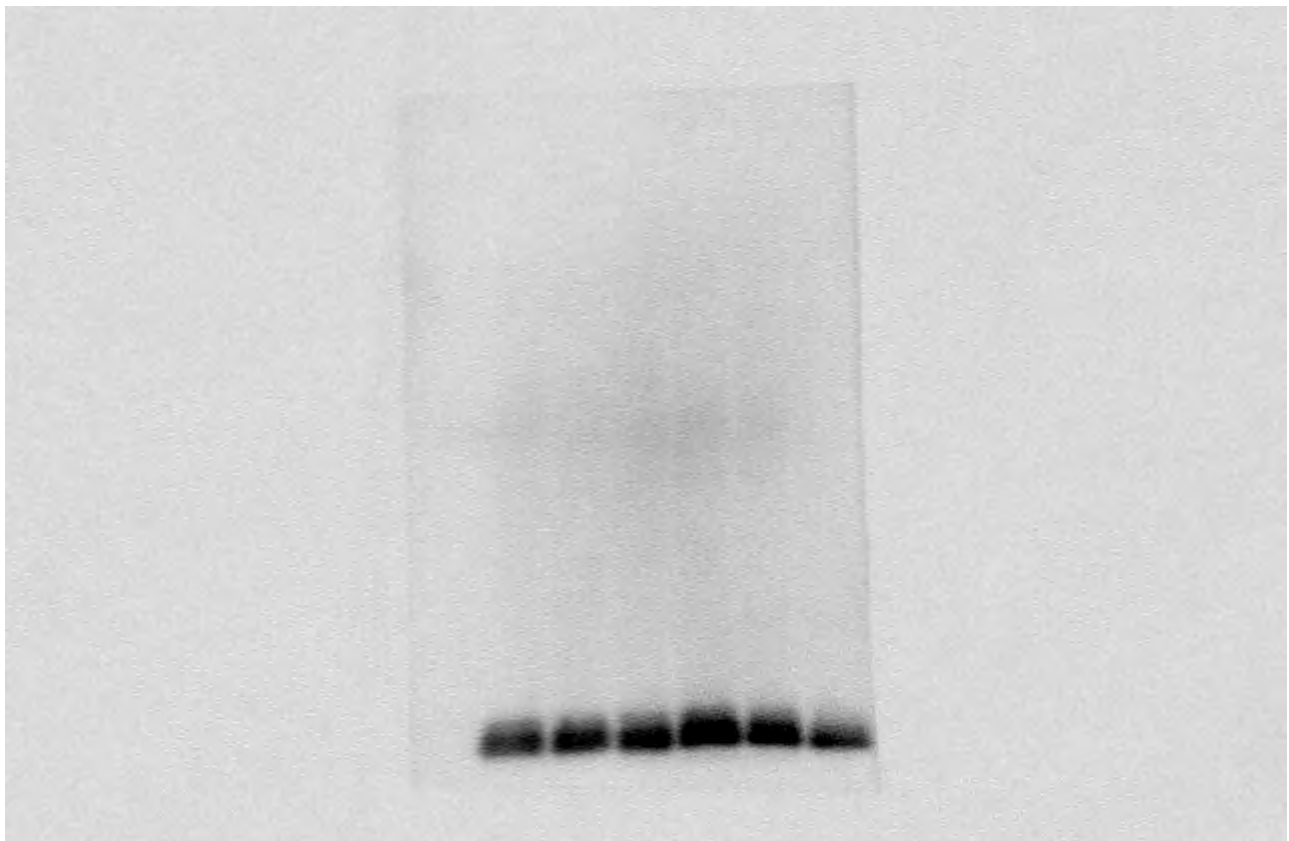


Figure S23 Western blot for 4EBP1 of A498 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

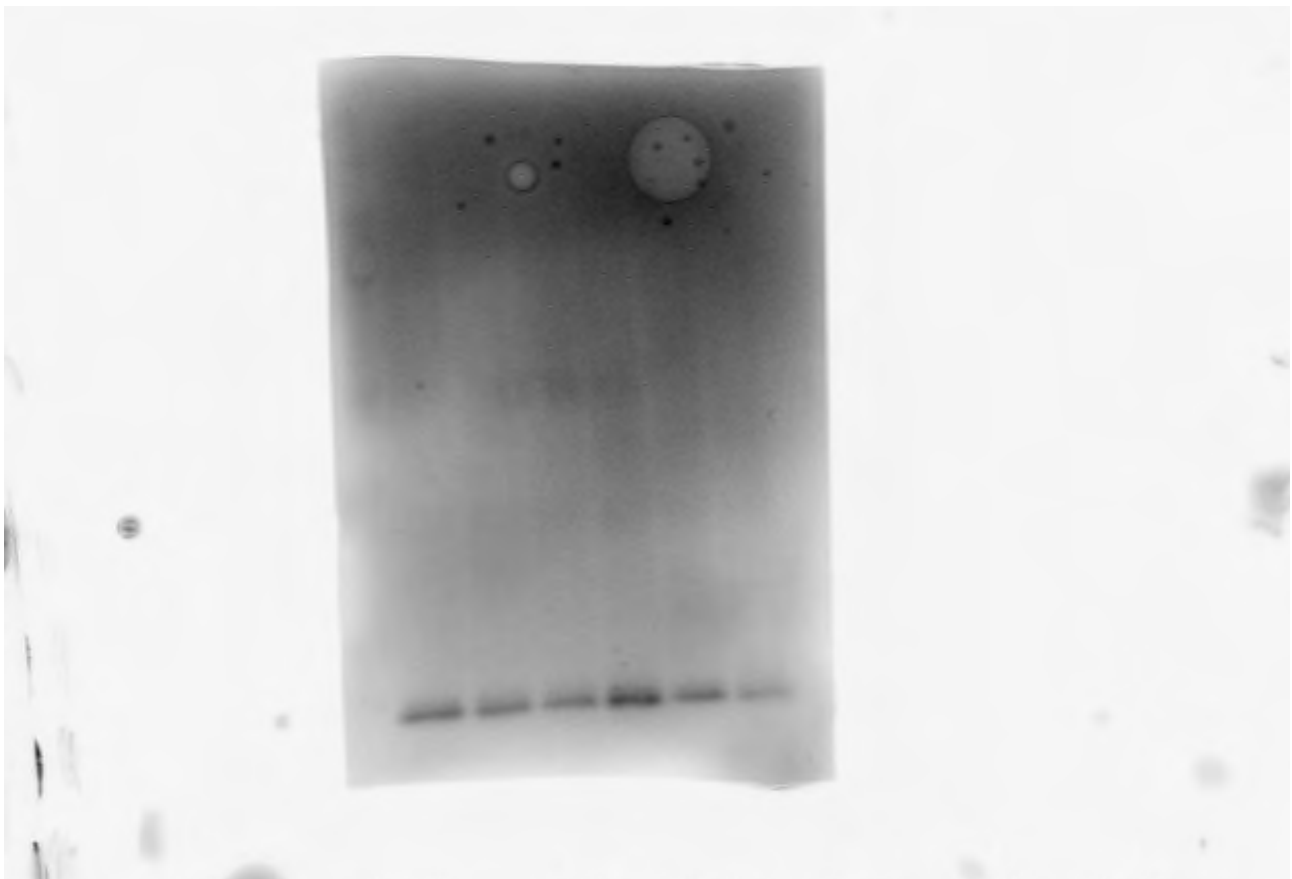


Figure S24 Western blot for phospho-4EBP1(Thr70) of A498 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

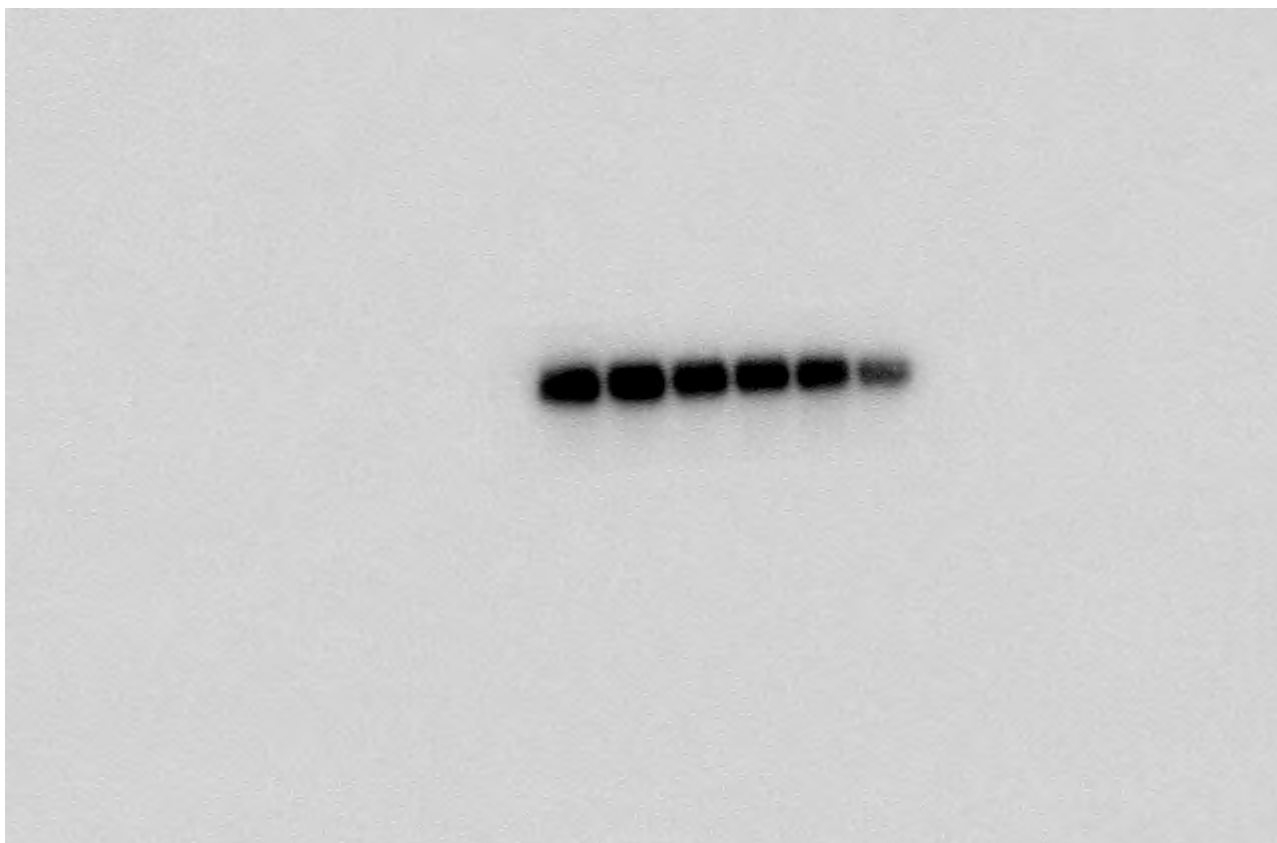


Figure S25 Western blot for S6RP of A498 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.



Figure S26 Western blot for phospho-S6RP(Ser235/236) of A498 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.



Figure S27 Western blot for ERK1/2 of A498 cell line after BLU9931 treatment (above). From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration. (The below membrane showed a reproductive result of phospho-ERK1/2.)

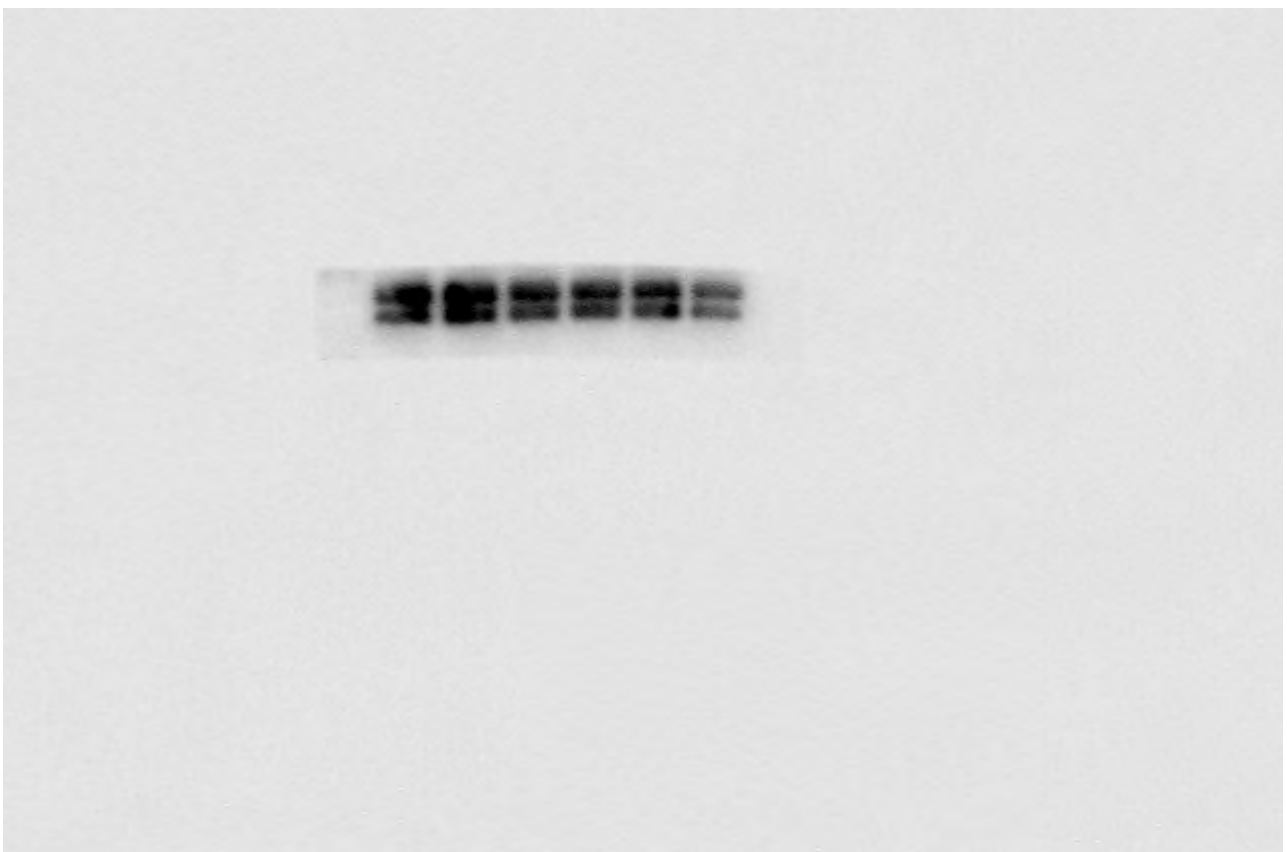


Figure S28 Western blot for phospho-ERK1/2(Thr202/Tyr204) of A498 cell line after BLU9931 treatment. From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

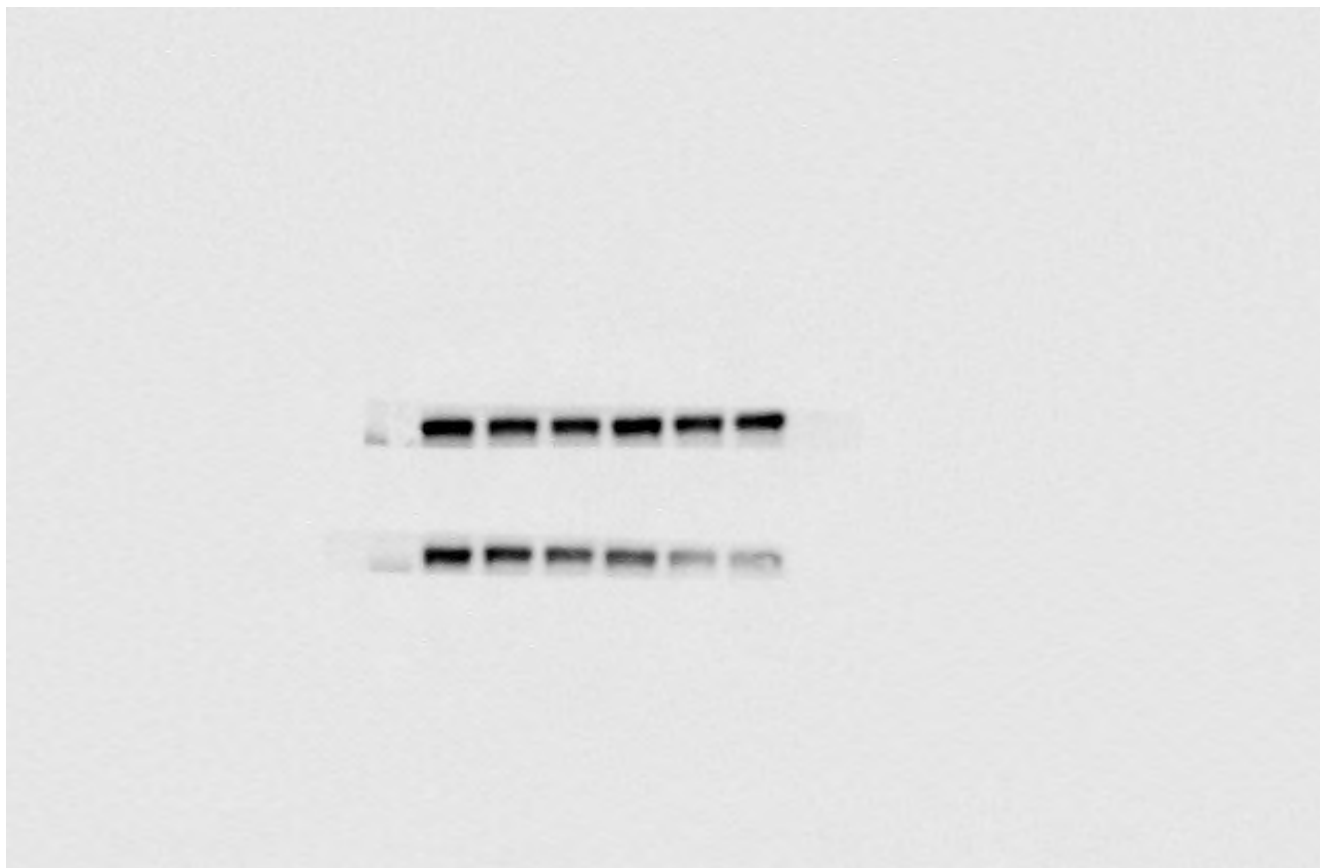


Figure S29 Western blot for STAT3(above) and phospho-STAT3(Tyr705, below) of A498 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

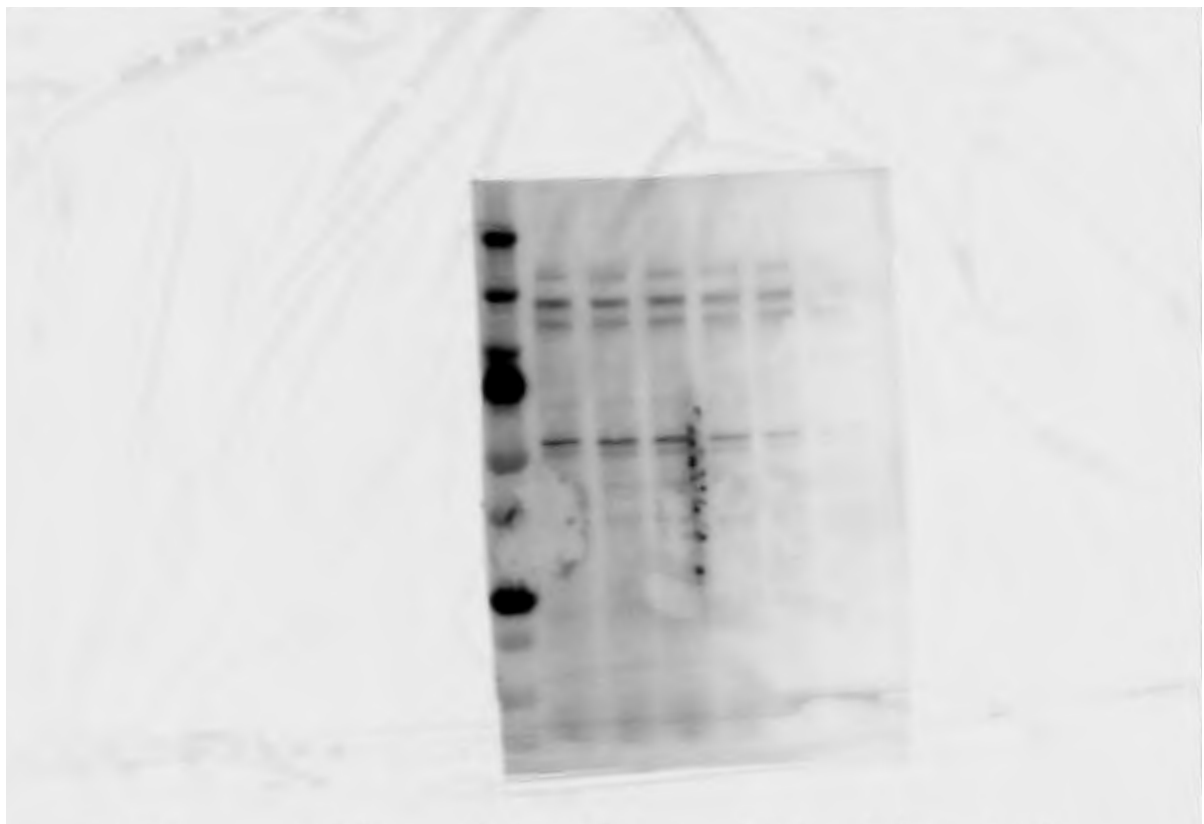


Figure S30 Western blot for phospho-FGFR(Thr653/654) of A704 cell line after BLU9931 treatment. From left, MW marker, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

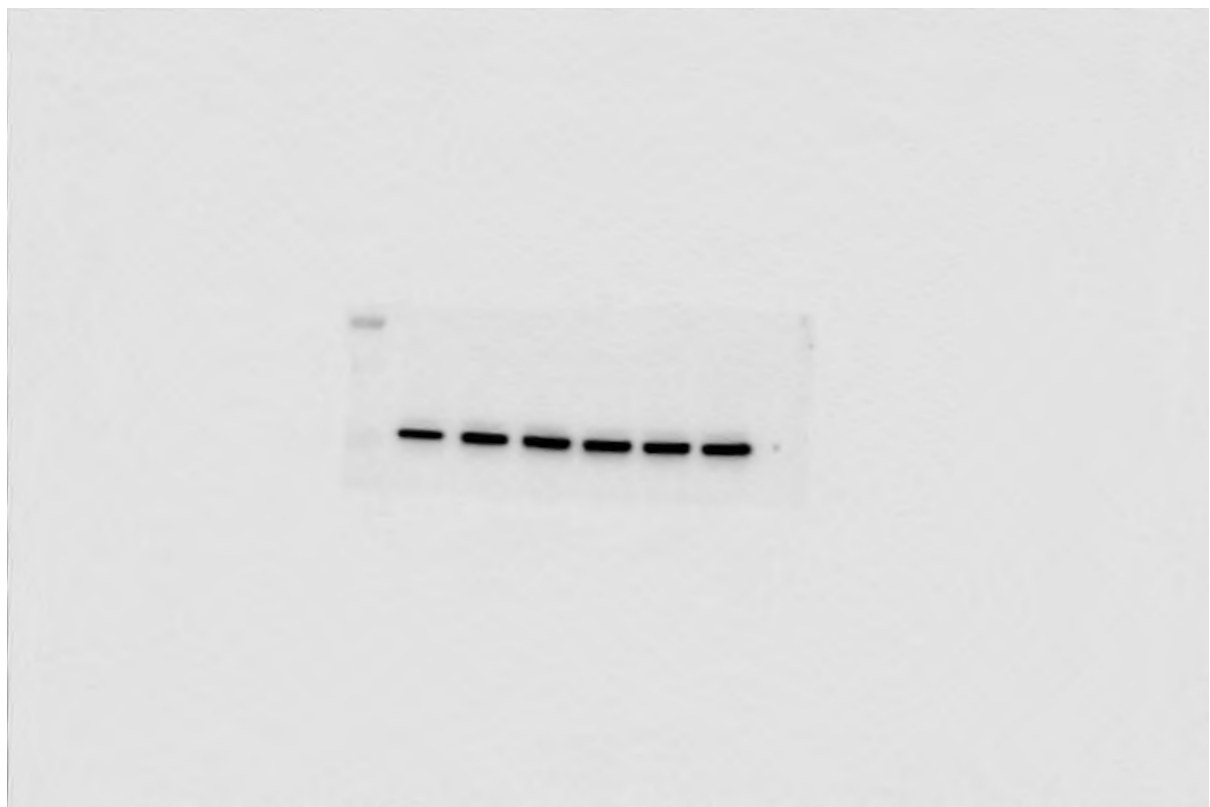


Figure S31 Western blot for α -Tubulin of A704 cell line after BLU9931 treatment. From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

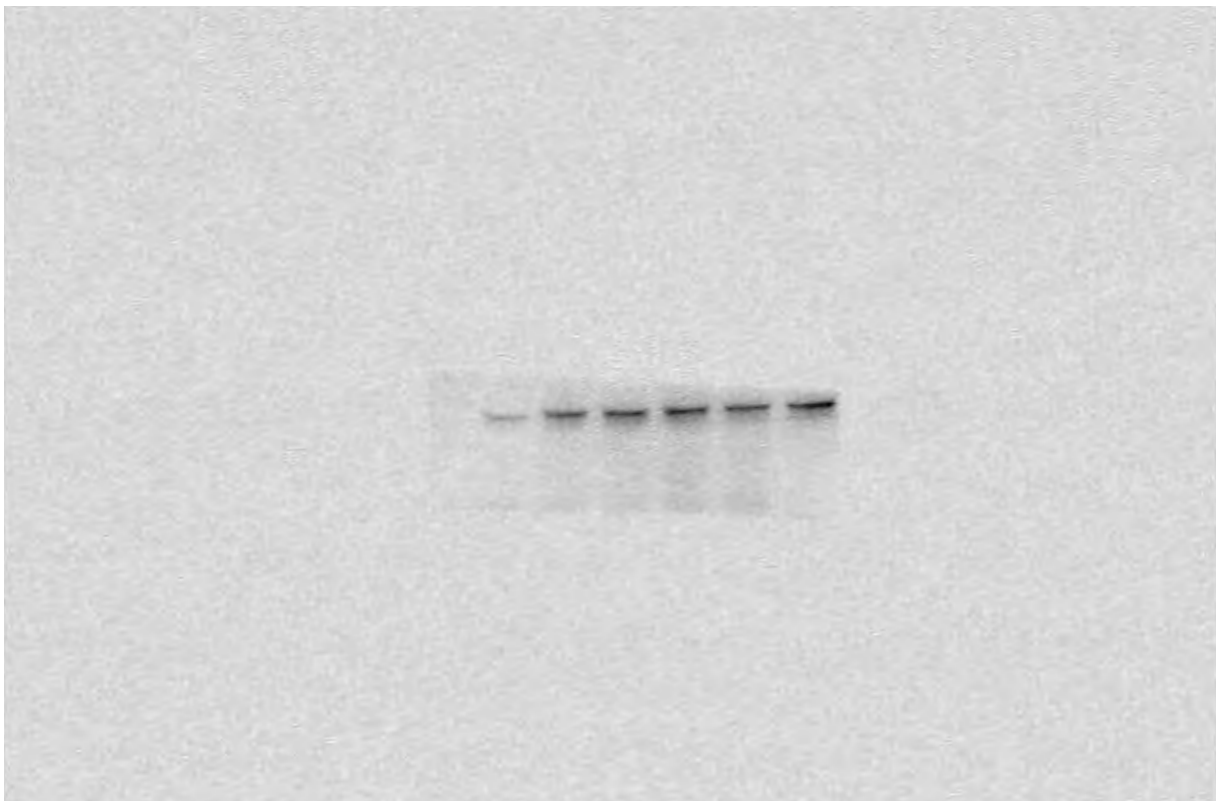


Figure S32 Western blot for AKT of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.



Figure S33 Western blot for phospho-AKT(Ser473) of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.



Figure S34 Western blot for 4EBP1 of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.



Figure S35 Western blot for phospho-4EBP1(Thr70) of A704 cell line after BLU9931 treatment. From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.



Figure S36 Western blot for S6RP of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

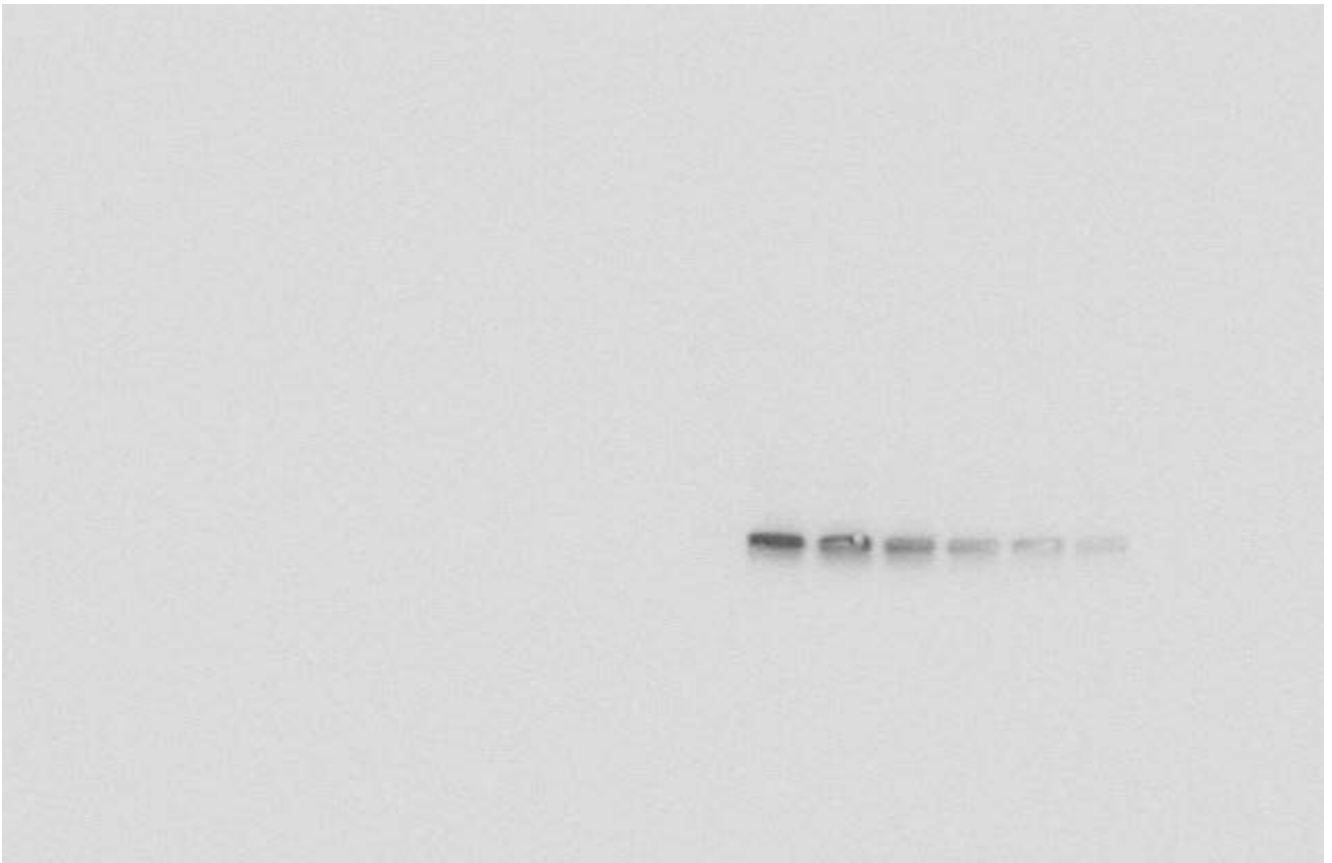


Figure S37 Western blot for phospho-S6RP(Ser235/236) of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

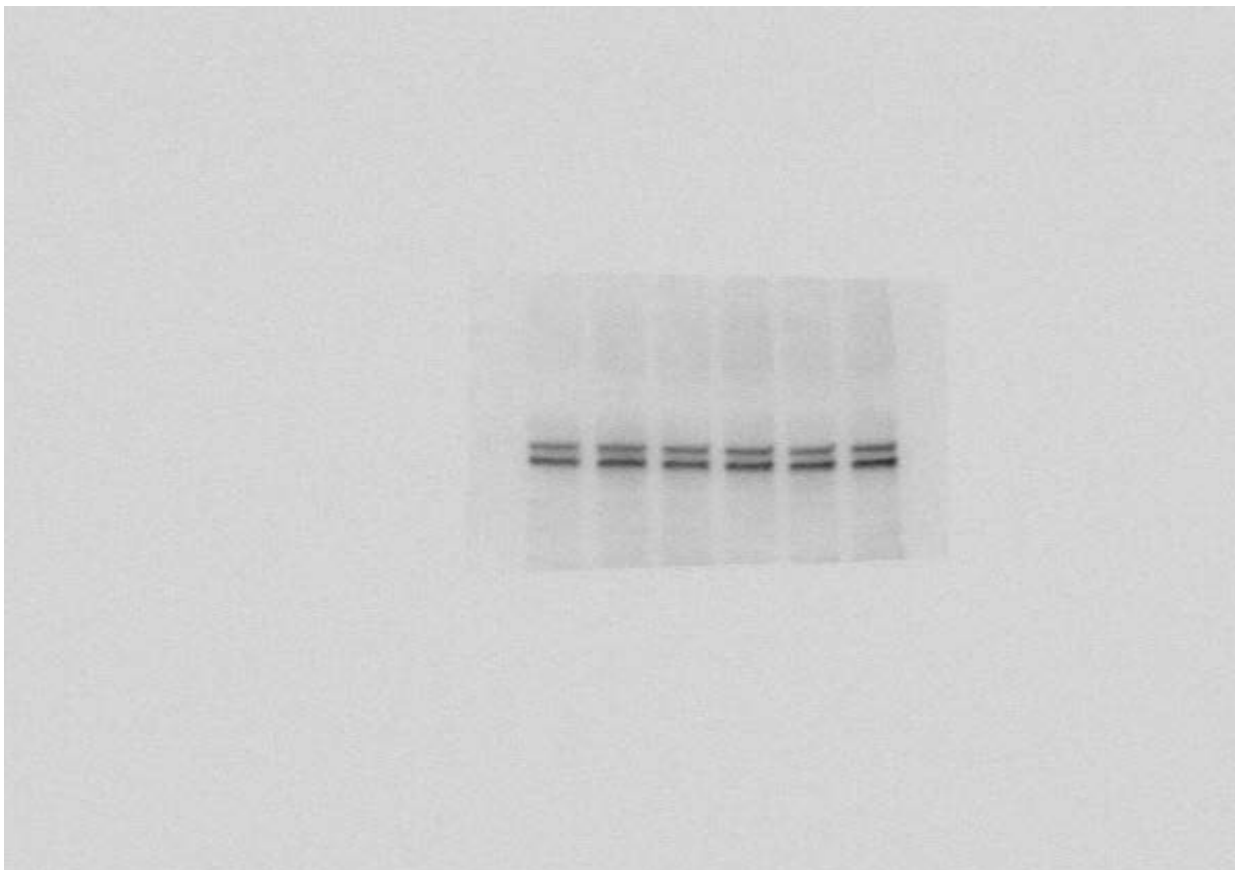


Figure S38 Western blot for ERK1/2 of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

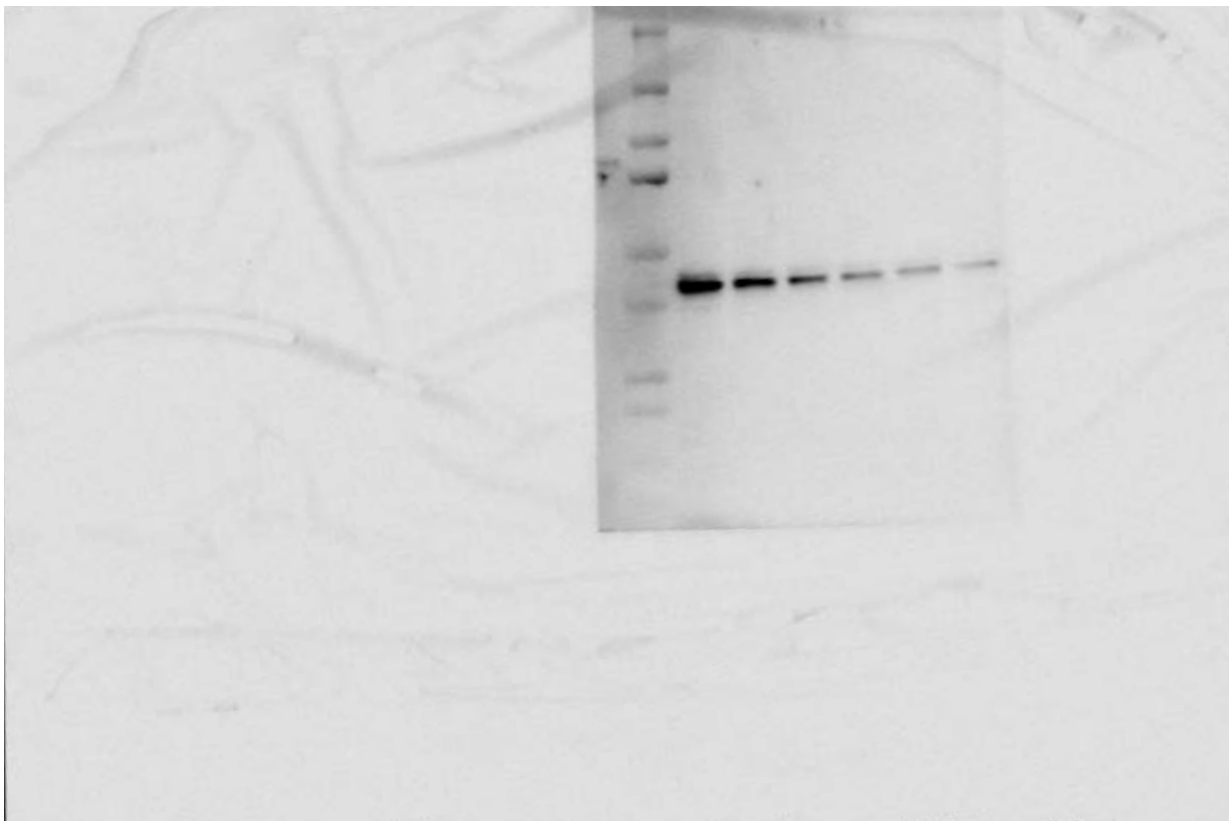


Figure S39 Western blot for phospho-ERK1/2(Thr202/Tyr204) of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

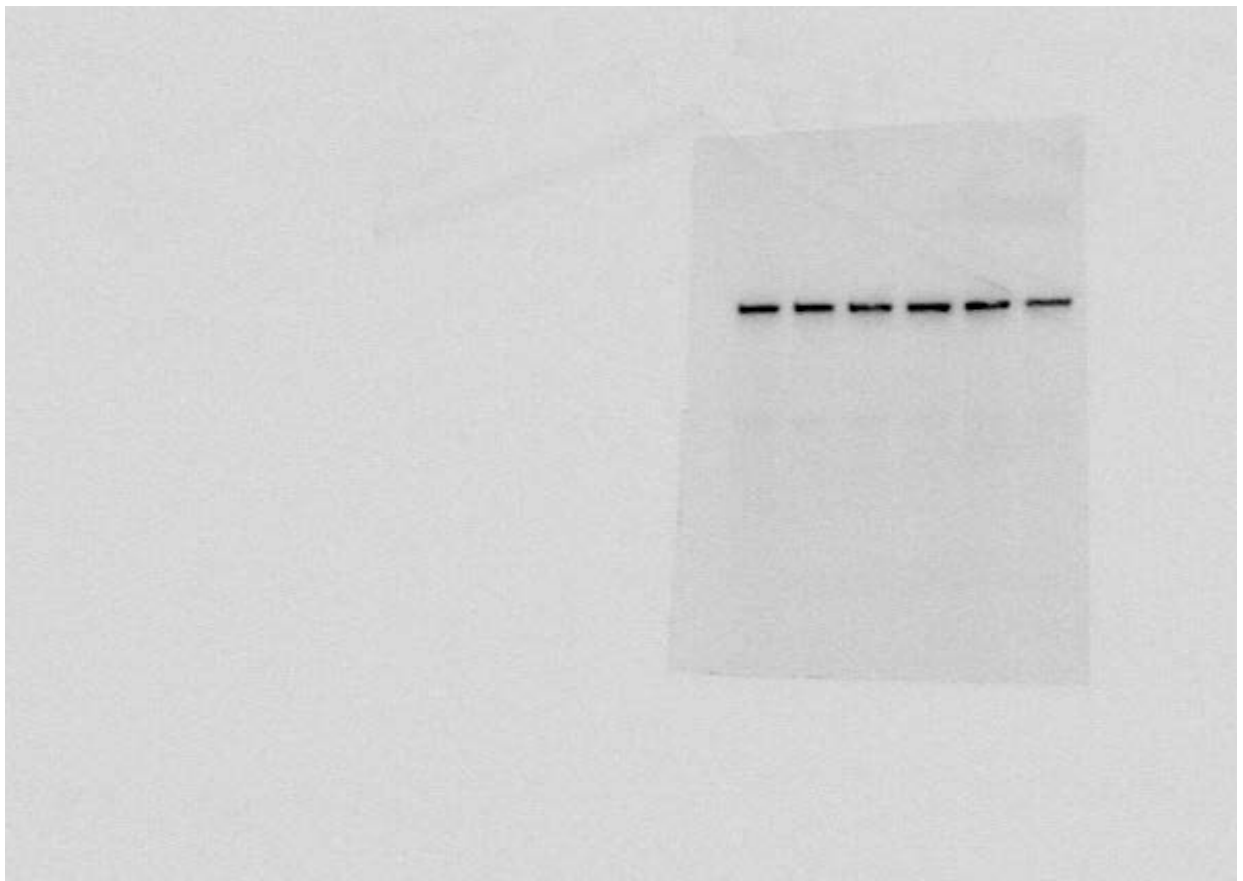


Figure S40 Western blot for STAT3 of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

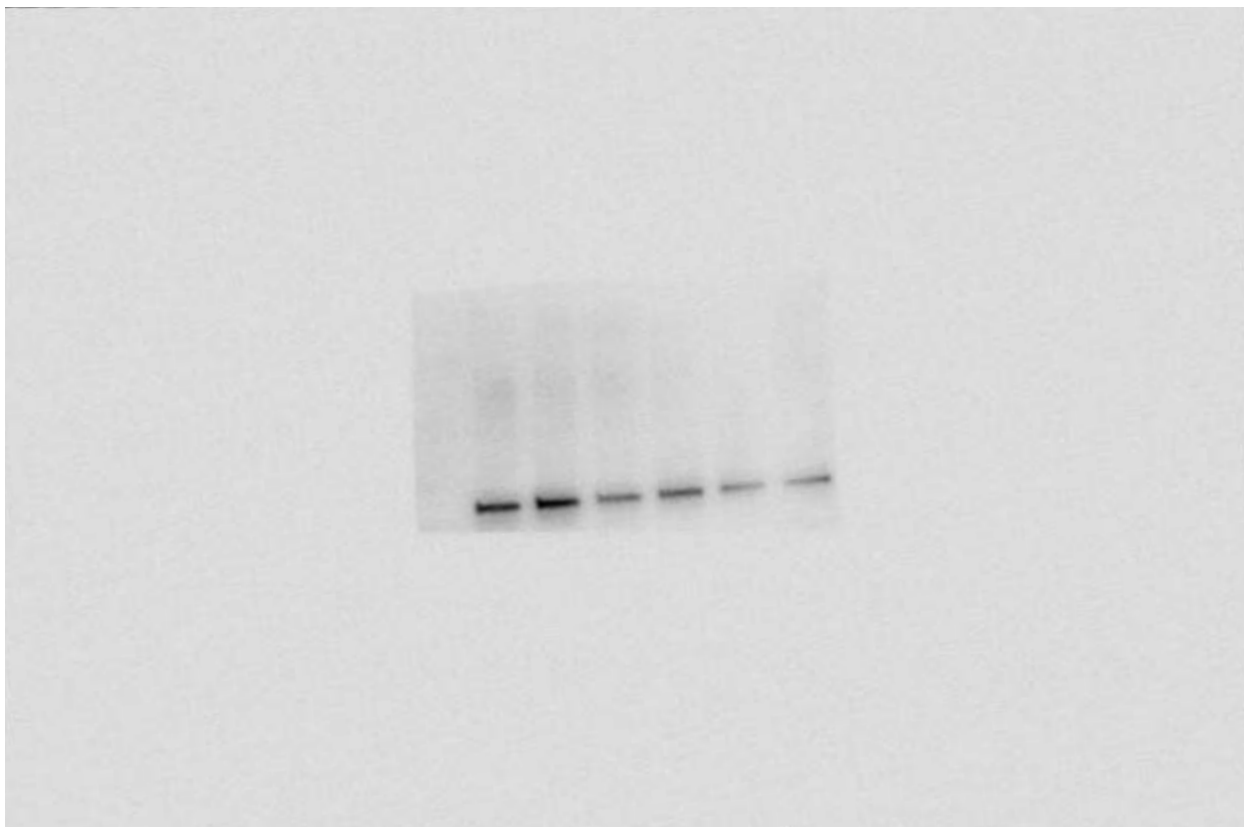


Figure S41 Western blot for phospho-STAT3(Tyr705) of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

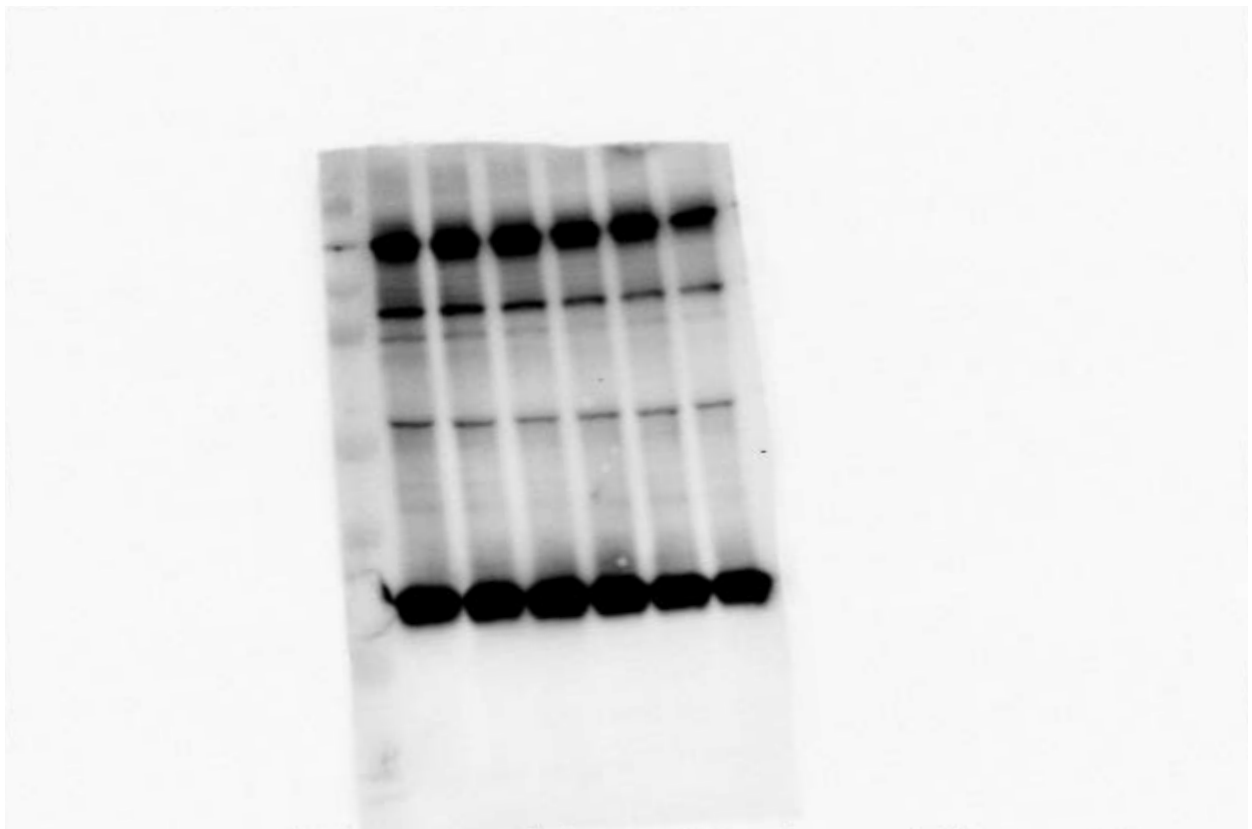


Figure S42 Western blot for phospho-FGFR(Thr653/654) of 769P cell line after BLU9931 treatment. From left, MW marker, Control, 1 nM, 10 nM, 100 nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

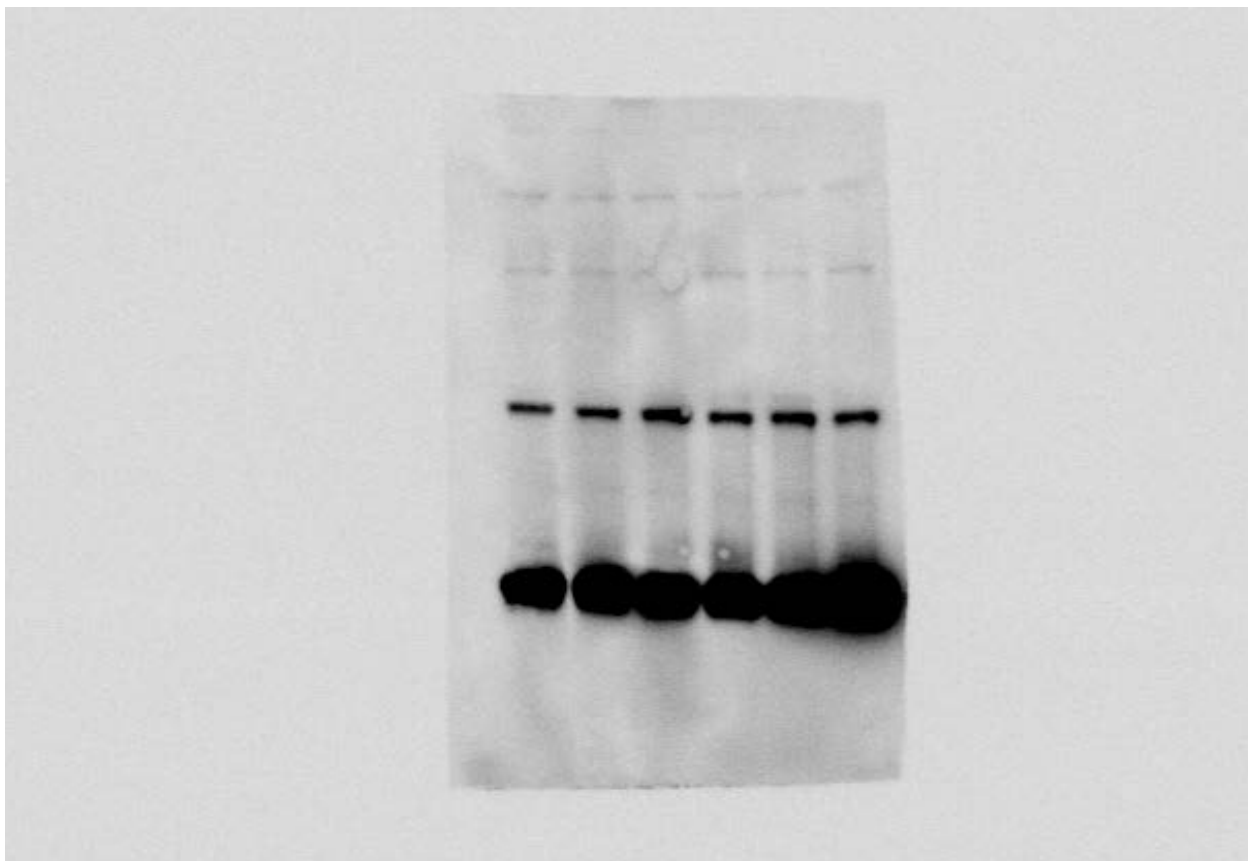


Figure S43 Western blot for α -Tubulin of 769P cell line after BLU9931 treatment. From left, Control, 1 nM, 10 nM, 100 nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

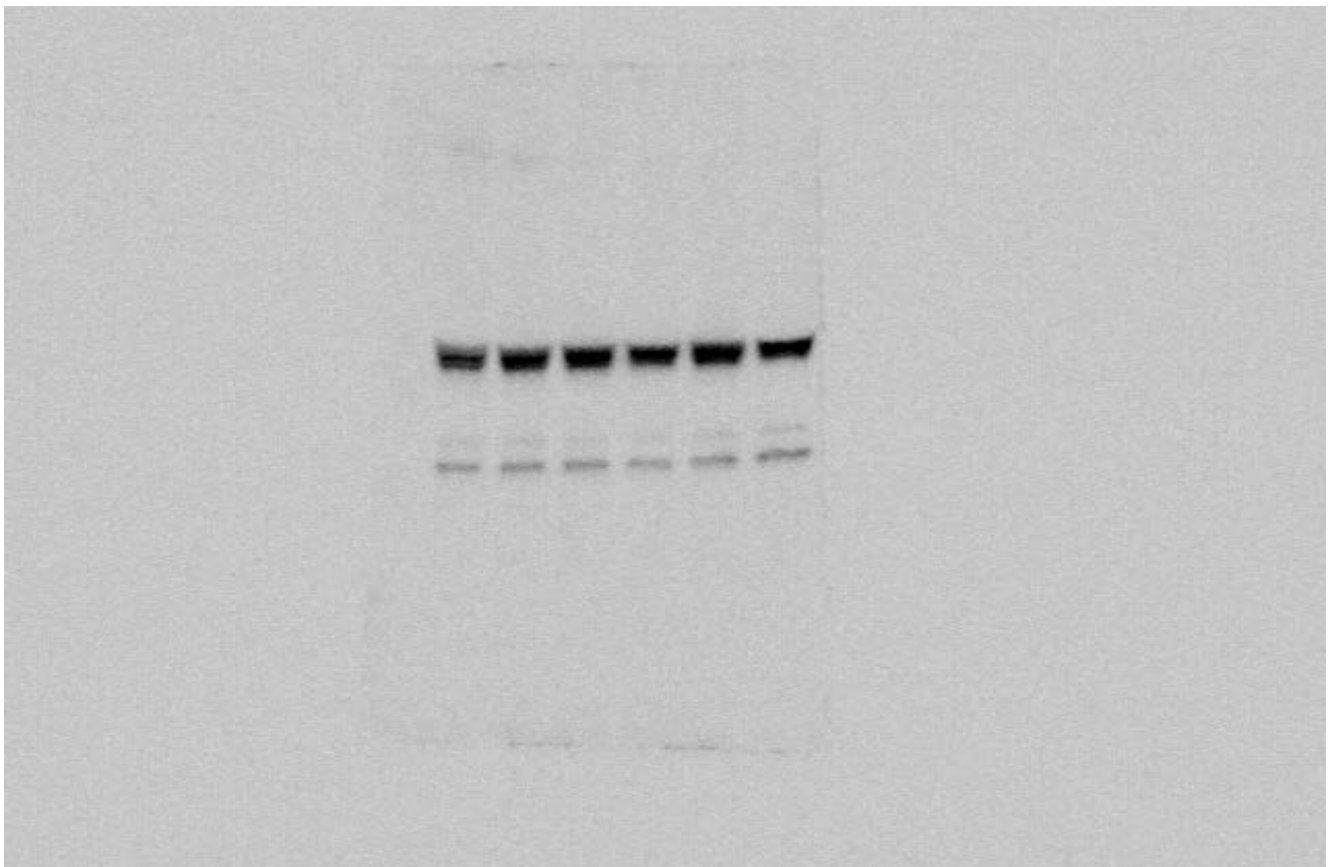


Figure S44 Western blot for AKT of 769P cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment.

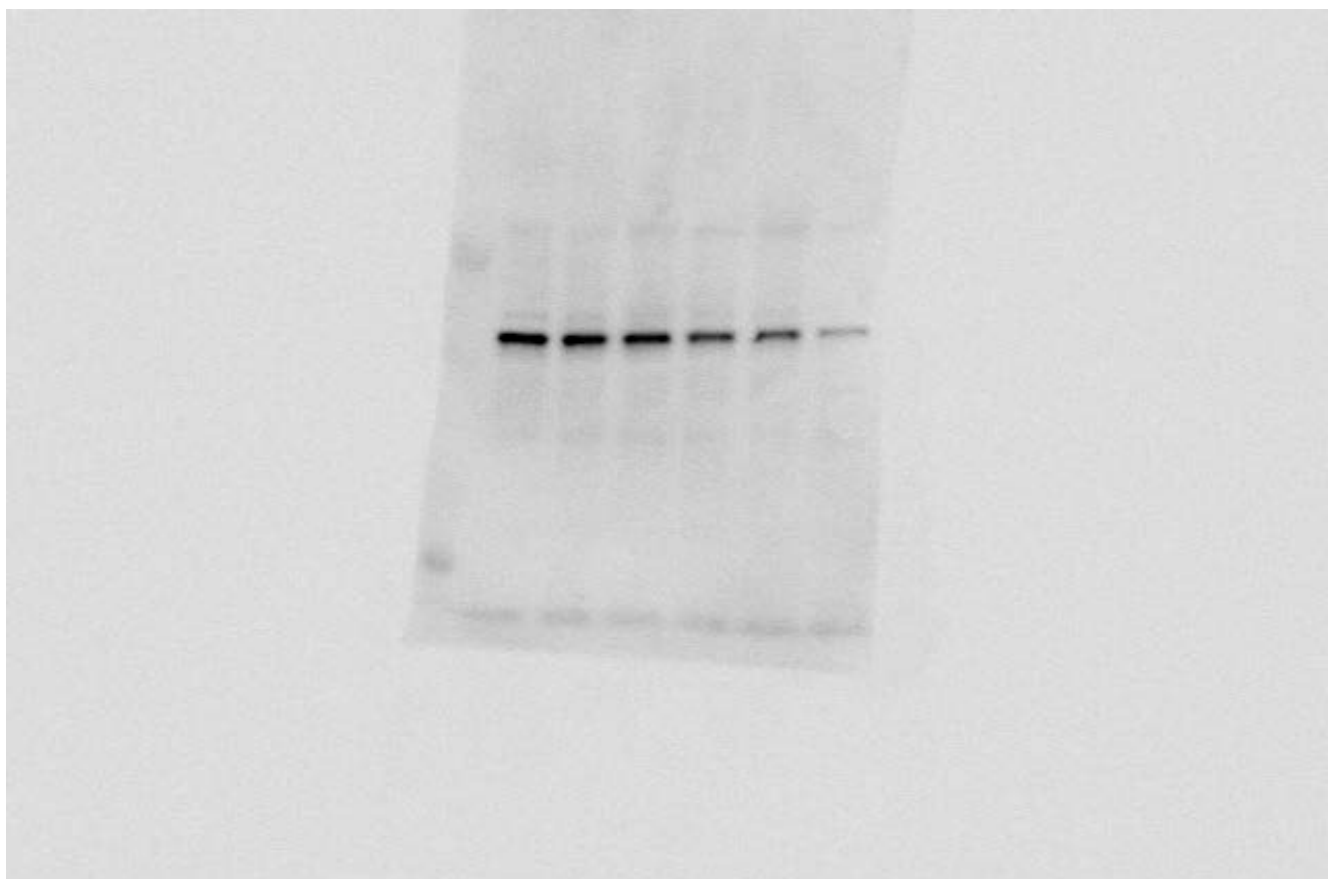


Figure S45 Phospho-AKT(Ser 473) of 769P cell line after BLU9931 treatment.

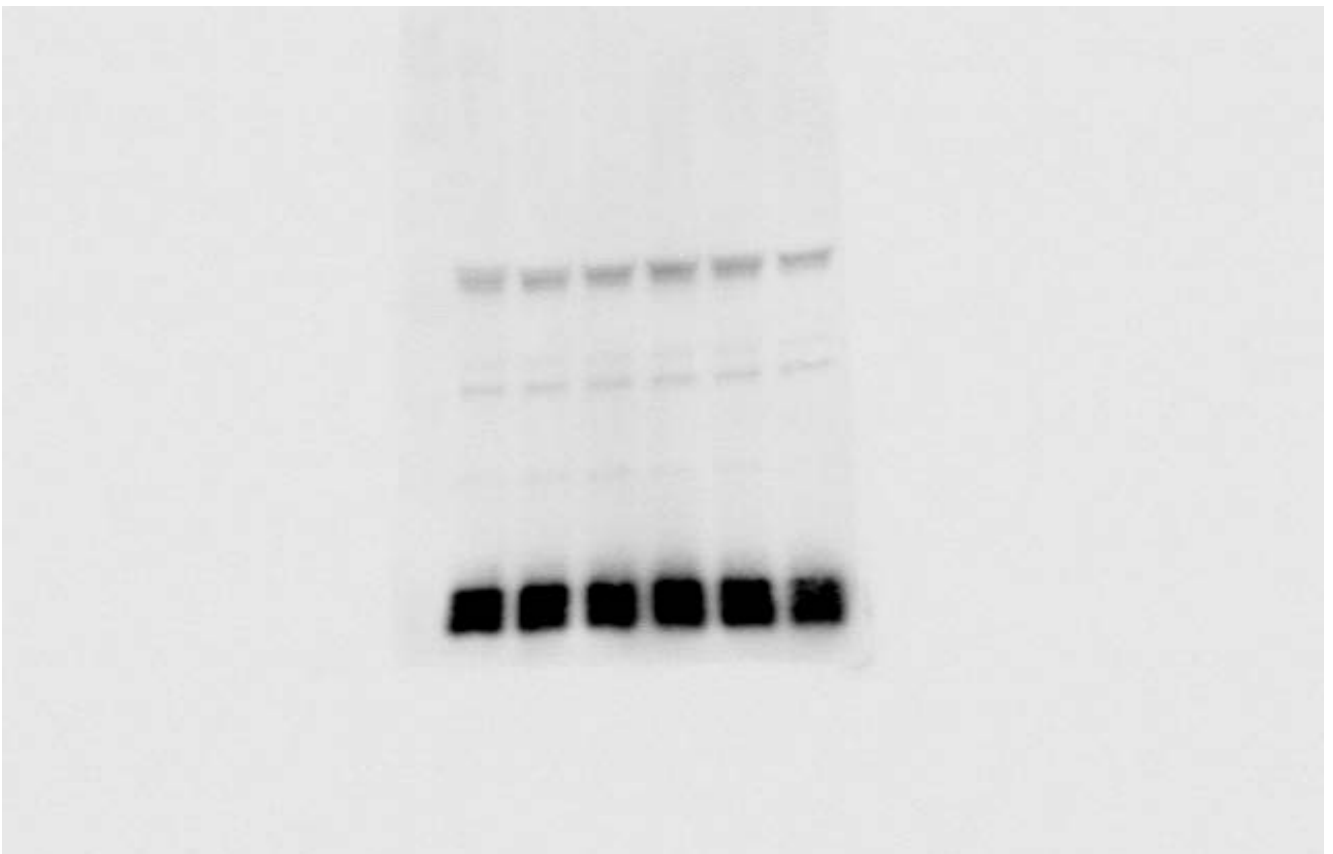


Figure S46 Western blot for 4EBP1 of 769P cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

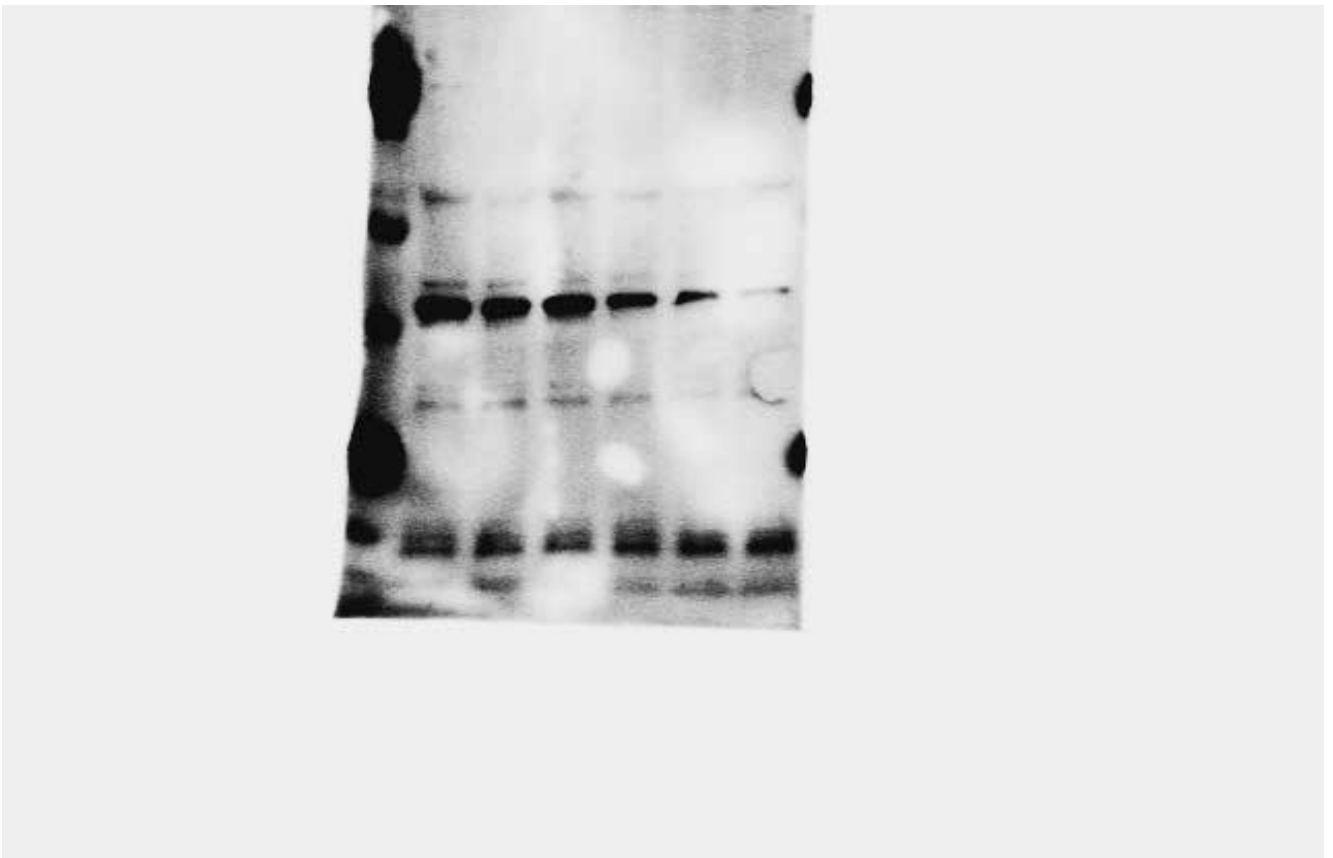


Figure S47 Western blot for phospho-4EBP1(Thr70) of 769P cell line after BLU9931 treatment. From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

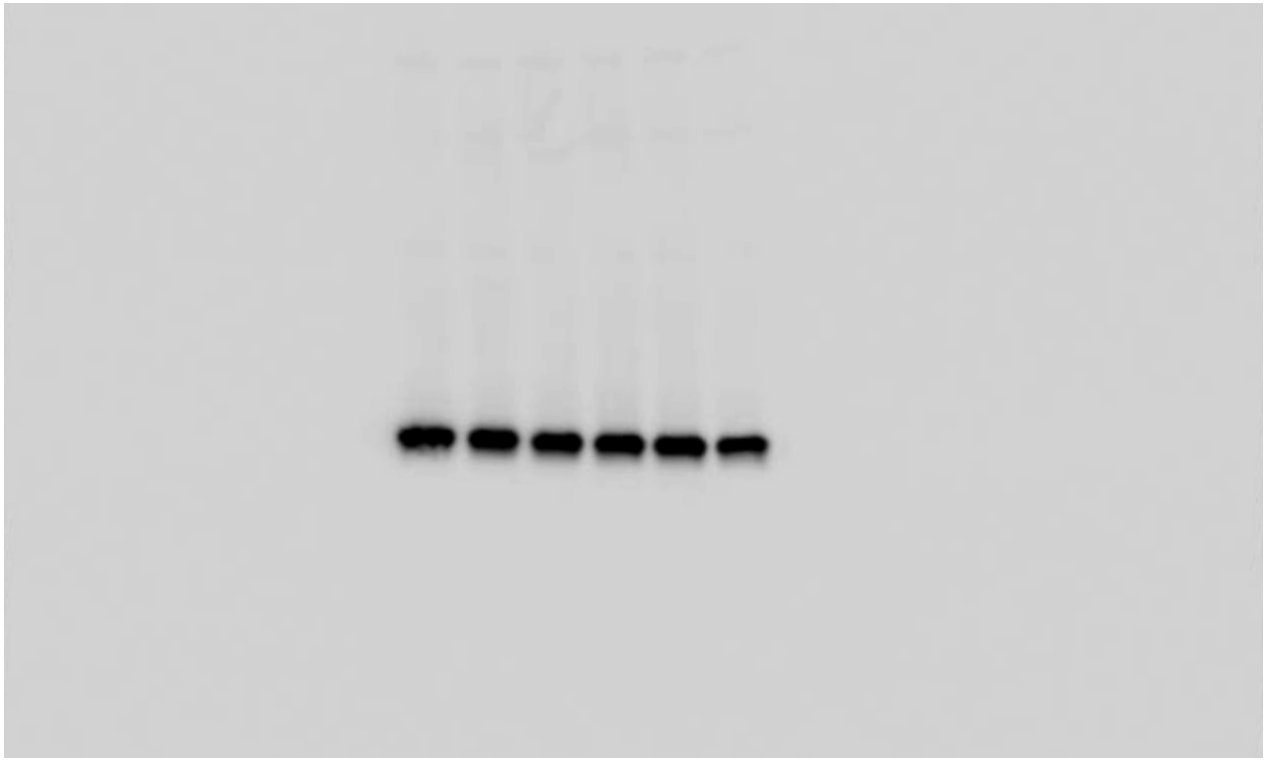


Figure S48 Western blot for S6RP of 769P cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

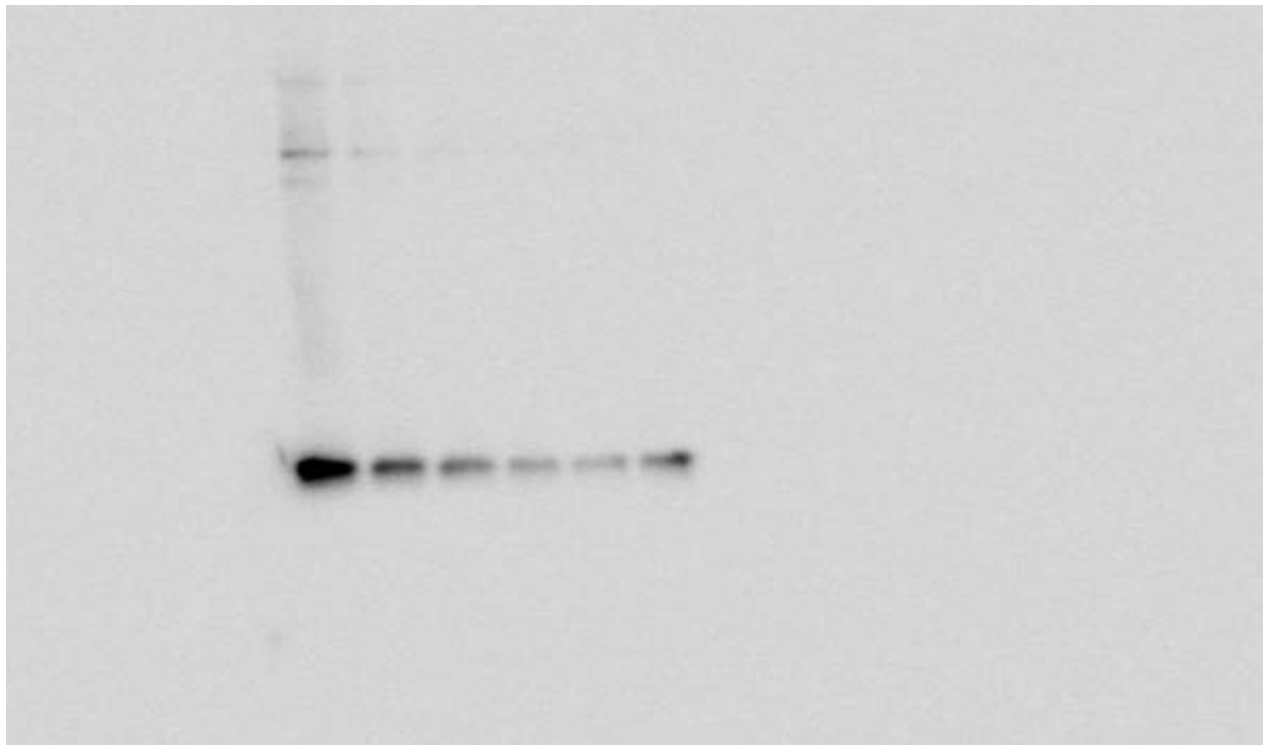


Figure S49 Western blot for phospho-S6RP(Ser235/236) of 769P cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

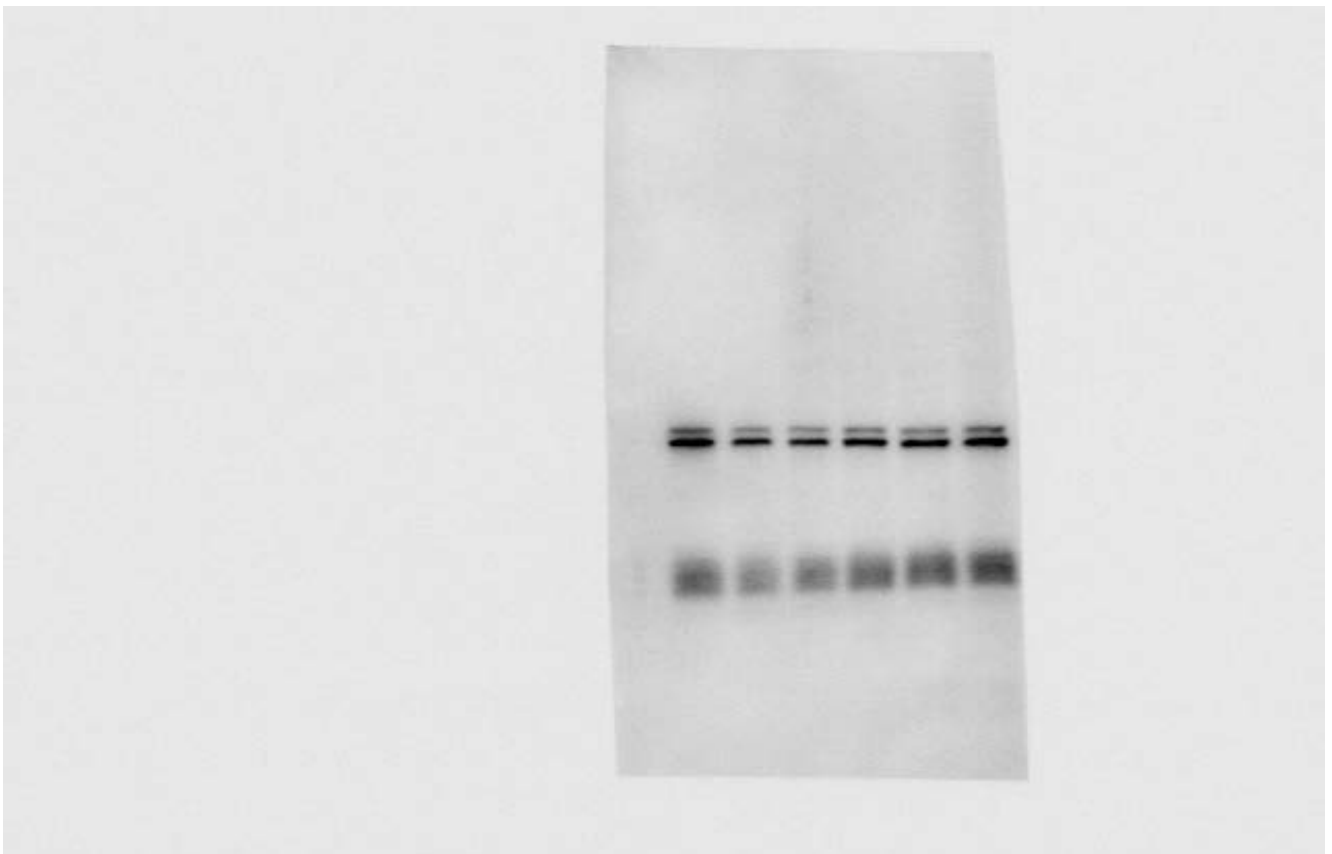


Figure S50 Western blot for ERK1/2 of 769P cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

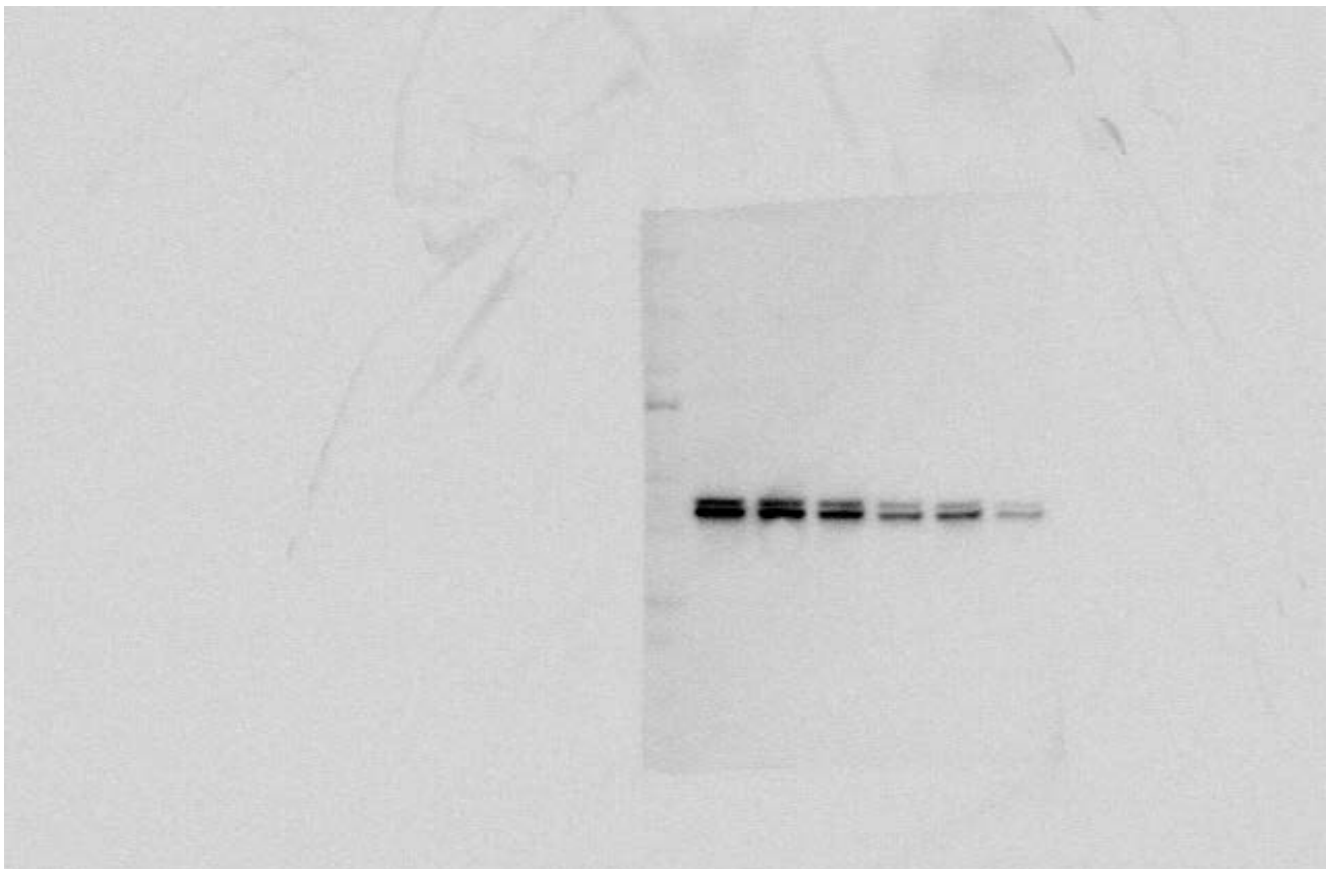


Figure S51 Western blot for phospho-ERK1/2(Thr202/Tyr204) of 769P cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

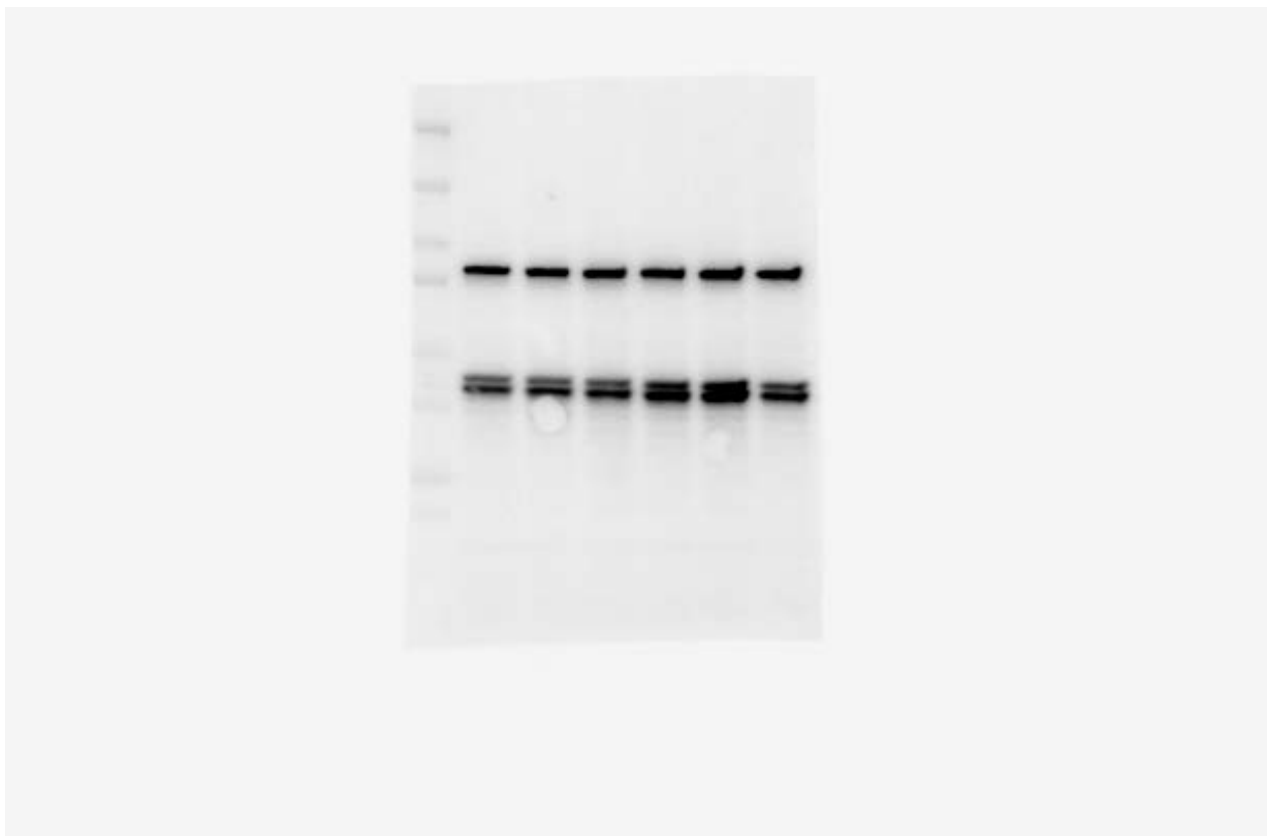


Figure S52 Western blot for STAT3 of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.

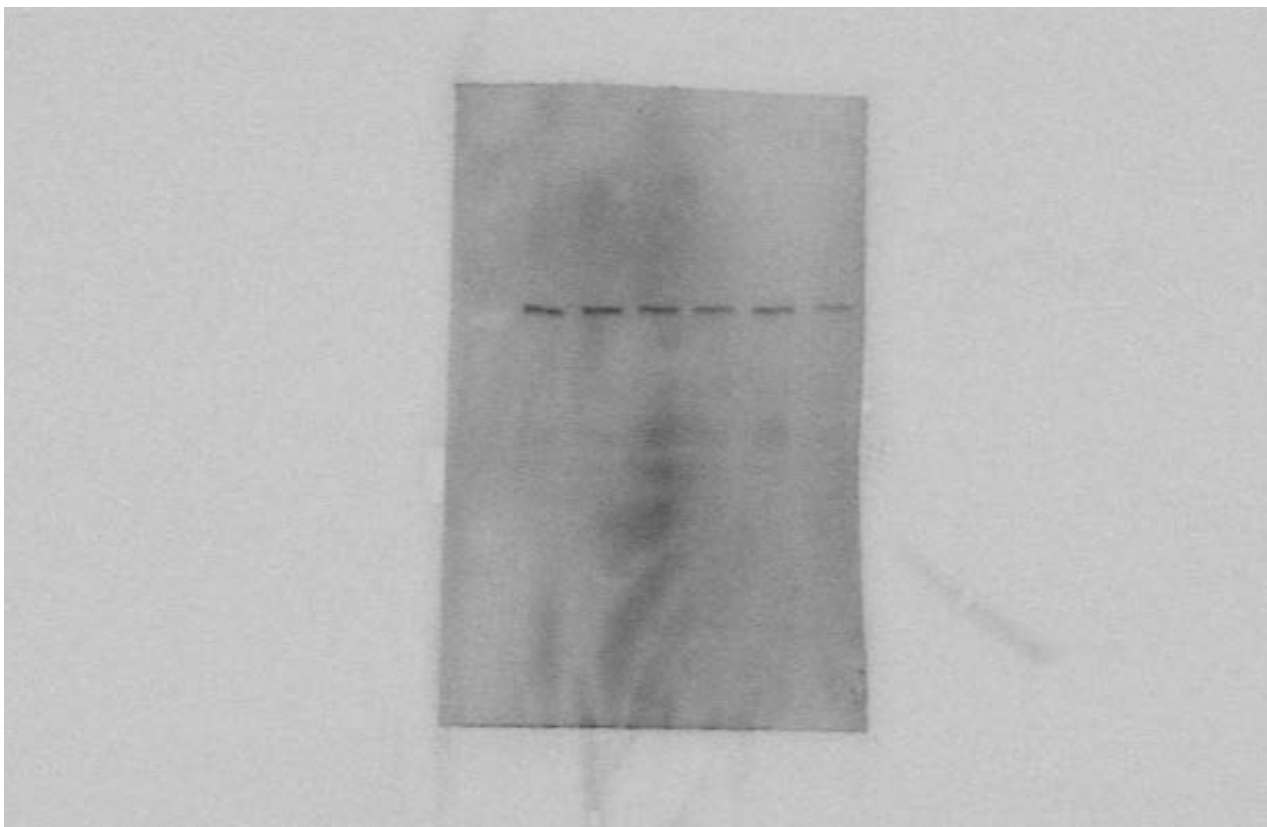


Figure S53 Western blot for phospho-STAT3(Tyr705) of A704 cell line after BLU9931 treatment.
From left, Control, 1nM, 10nM, 100nM, 1 μ M, 10 μ M of BLU9931 treatment concentration.