Supplementary Table 1: Background of the patients of screening cohort

Background	PR group ¹	PD group ¹	P value ²	
	(n = 8)	(n=8)		
Age (years old)	68.5 (60.75 – 75.5)	73 (62.75 – 80.75)	0.2473	
Sex: male/female	7 / 1	5/3	0.5692	
Virus: HBV/HCV/NBNC	5 / 1 / 2	2/2/4	0.3189	
Maximum tumor size (cm)	4.7 (3.85 – 6.7)	4.2 (2.4 – 11.7)	1	
Number of tumor: $\leq 5 / \geq 6$	5/3	3 / 5	0.6193	
BCLC: B / C ³	5/3	1 / 7	0.1189	
Extrahepatic metastasis: without / with	2 / 6	4 / 4	0.6084	
Vascular invasion:	7 / 1	5 / 3	0.5692	
without / with				
Initial dose of sorafenib: 800mg / 400mg	6 / 2	6 / 2	1	
AFP before treatment (ng/ml)	201 (12 – 1644)	4009.5 (9 – 12345)	0.4622	
DCP before treatment (mAU/ml)	212 (28.25 – 1785)	6533.5 (2173 – 380064)	0.0157	
Δ AFP 4	-0.47 (-0.750.19)	0.55 (-0.03 – 1.19)	0.0074	
ΔDCP ⁴	0.13 (-0.38 – 15.6)	3.03 (1.45 – 5.64)	0.3446	

¹Median value (25th–75th percentiles) for contentious variables and number of the patients for categorical variables are shown.

 $^{^{2}}$ *P* values by the Wilcoxon rank-sum test for contentious variables and by the Fischer's exact test for categorical variables are shown. The bold represent the *p* value of <0.05.

³ There is no HCC patient with BCLC stage A.

⁴ (Serum level after treatment – serum level before treatment)/ serum level before

treatment. The minus value represents the decrease of tumor marker after the sorafenib treatment.

PR: the group showing partial response at 3 months after the initiation of treatment.

PD: the group showing progressive disease at one month after the initiation of treatment.

Supplementary Table 2: Clinical factors and serum miR-181a-5p level for affecting overall survival in patients with BCLC-C HCC.

variables	Number of patients	P value ¹ (univariate)	P value ² (multivariate)	Hazard ratio (95% CI)
Age (yeas old)				
$73 \geq$	17	0.1204	_	_
74 <u>≤</u>	13	0.120		
Sex				
male	24	0.1738	-	-
female	6			
Maximum size				
$3.6 \text{ cm} \ge$	13	0.2666	-	-
$3.7 \text{ cm} \leq$	17			
Number of tumor				
5 <u>≥</u>	12	0.0194	0.1549	0.491 (0.165-1.30)
6 ≤	18			1
Extrahepatic				
metastasis				
presence	23	0.9519	-	-
absence	7			
Portal vein				
thrombosis				
presence	11	0.6412	-	-
absence	19			
Serum miR-181a-5p level				
high	10	0.0037	0.0194	0.267 (0.070-0.818)
low	20	0.0007	V•V1/T	1

¹ P value by log-rank test.

Continuous valuables of age, tumor size and number of tumor were categorized using the median values. Bold denotes p < 0.05.

 $^{^2}P$ value by Cox proportional-hazards regression model.