

Table S1. List of the pre-S genotyping results by TA cloning- and NGS-based analyses in 75 HBV-related HCC patients

Patient No.	TA Cloning Result ^a	NGS Result (Pre-S Del Type (%)) ^b	NGS Result (Pre-S Del Region (%)) ^c
1	pre-S1 del (nt 2855-2872)	1. pre-S1 del (92.118)^d 2. wild-type (7.278) 3. pre-S2 del (0.372) 4. pre-S1+pre-S2 del (0.231) I. pre-S1 plus pre-S2 del (92.490) ^e II. pre-S1 plus pre-S1+pre-S2 del (92.349) III. pre-S2 plus pre-S1+pre-S2 del (0.603) IV. all three types of pre-S del (92.722)	1. pre-S1 del (nt 2855-2872) (90.102) 2. wild-type (7.278) 3. pre-S2 del (nt 1-28) (0.180) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-111) (0.027)
2	1. wild-type 2. pre-S1 del (nt 2910-3089)	1. pre-S1 del (75.241) 2. wild-type (22.338) 3. pre-S2 del (1.891) 4. pre-S1+pre-S2 del (0.530) I. pre-S1 plus pre-S2 del (77.132) II. pre-S1 plus pre-S1+pre-S2 del (75.771) III. pre-S2 plus pre-S1+pre-S2 del (2.421) IV. all three types of pre-S del (77.662)	1. pre-S1 del (nt 2910-3089) (48.977) 2. wild-type (22.338) 3. pre-S2 del (nt 1-54) (1.579) 4. pre-S1+pre-S2 del (nt 2910-3089, 1-57) (0.277)
3	wild-type	1. pre-S1 del (76.228) 2. wild-type (12.583) 3. pre-S2 del (10.622) 4. pre-S1+pre-S2 del (0.567) I. pre-S1 plus pre-S2 del (86.850) II. pre-S1 plus pre-S1+pre-S2 del (76.795) III. pre-S2 plus pre-S1+pre-S2 del (11.189) IV. all three types of pre-S del (87.417)	1. pre-S1 del (nt 3110-3127) (24.869) 2. wild-type (12.583) 3. pre-S2 del (nt 1-57) (10.000) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-57) (0.490)

4	wild-type	1. wild-type (71.590) 2. pre-S1 del (28.270) 3. pre-S2 del (0.129) 4. pre-S1+pre-S2 del (0.011) I. pre-S1 plus pre-S2 del (28.399) II. pre-S1 plus pre-S1+pre-S2 del (28.281) III. pre-S2 plus pre-S1+pre-S2 del (0.140) IV. all three types of pre-S del (28.410)	1. wild-type (71.590) 2. pre-S1 del (nt 2854-3147) (23.961) 3. pre-S2 del (nt 3211-3216) (0.048) 4. pre-S1+pre-S2 del (nt 2954-3097, 1-54) (0.002)
5	pre-S2 del (nt 24-50)	1. wild-type (58.461) 2. pre-S2 del (37.194) 3. pre-S1 del (2.939) 4. pre-S1+pre-S2 del (1.407) I. pre-S1 plus pre-S2 del (40.133) II. pre-S1 plus pre-S1+pre-S2 del (4.346) III. pre-S2 plus pre-S1+pre-S2 del (38.601) IV. all three types of pre-S del (41.539)	1. wild-type (58.461) 2. pre-S2 del (nt 24-50) (33.801) 3. pre-S1 del (nt 2880-3146) (0.646) 4. pre-S1+pre-S2 del (nt 2880-3146, 24-50) (0.355)
6	wild-type	1. wild-type (96.592) 2. pre-S1 del (2.906) 3. pre-S2 del (0.469) 4. pre-S1+pre-S2 del (0.033) I. pre-S1 plus pre-S2 del (3.375) II. pre-S1 plus pre-S1+pre-S2 del (2.939) III. pre-S2 plus pre-S1+pre-S2 del (0.502) IV. all three types of pre-S del (3.408)	1. wild-type (96.592) 2. pre-S1 del (nt 3026-3205) (0.604) 3. pre-S2 del (nt 1-12) (0.233) 4. pre-S1+pre-S2 del (nt 2854-2979, 6-134) (0.008)
7	wild-type	1. wild-type (99.000) 2. pre-S1 del (0.927)	1. wild-type (99.000) 2. pre-S1 del (nt 3106-3129) (0.294)

		3. pre-S2 del (0.062) 4. pre-S1+pre-S2 del (0.012) I. pre-S1 plus pre-S2 del (0.989) II. pre-S1 plus pre-S1+pre-S2 del (0.939) III. pre-S2 plus pre-S1+pre-S2 del (0.074) IV. all three types of pre-S del (1.000)	3. pre-S2 del (nt 1-54) (0.031) 4. pre-S1+pre-S2 del (nt 2855-2972, 1-54) (0.008)
8	wild-type	1. wild-type (93.746) 2. pre-S1 del (5.846) 3. pre-S2 del (0.389) 4. pre-S1+pre-S2 del (0.018) I. pre-S1 plus pre-S2 del (6.235) II. pre-S1 plus pre-S1+pre-S2 del (5.864) III. pre-S2 plus pre-S1+pre-S2 del (0.407) IV. all three types of pre-S del (6.254)	1. wild-type (93.746) 2. pre-S1 del (nt 3026-3205) (2.417) 3. pre-S2 del (nt 3211-3213) (0.135) 4. pre-S1+pre-S2 del (nt 2856-2873, 1-21) (0.004)
9	wild-type	1. pre-S1+pre-S2 del (46.237) 2. pre-S2 del (26.927) 3. pre-S1 del (14.368) 4. wild-type (12.467) I. pre-S1 plus pre-S2 del (41.295) II. pre-S1 plus pre-S1+pre-S2 del (60.605) III. pre-S2 plus pre-S1+pre-S2 del (73.164) IV. all three types of pre-S del (87.533)	1. pre-S1+pre-S2 del (nt 2956-3126, 1-9) (24.002) 2. pre-S2 del (nt 1-54) (23.226) 3. pre-S1 del (nt 2944-3075) (5.140) 4. wild-type (12.467)
10	wild-type	1. wild-type (97.851) 2. pre-S1 del (2.037) 3. pre-S2 del (0.108) 4. pre-S1+pre-S2 del (0.005)	1. wild-type (97.851) 2. pre-S1 del (nt 3026-3205) (1.109) 3. pre-S2 del (nt 3211-3213) (0.049) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.003)

		I. pre-S1 plus pre-S2 del (2.145) II. pre-S1 plus pre-S1+pre-S2 del (2.042) III. pre-S2 plus pre-S1+pre-S2 del (0.113) IV. all three types of pre-S del (2.149)	
11	wild-type	1. wild-type (98.964) 2. pre-S1 del (0.943) 3. pre-S2 del (0.084) 4. pre-S1+pre-S2 del (0.009) I. pre-S1 plus pre-S2 del (1.027) II. pre-S1 plus pre-S1+pre-S2 del (0.952) III. pre-S2 plus pre-S1+pre-S2 del (0.093) IV. all three types of pre-S del (1.036)	1. wild-type (98.964) 2. pre-S1 del (nt 2854-2970) (0.125) 3. pre-S2 del (nt 1-9) (0.028) 4. pre-S1+pre-S2 del (nt 2954-3097, 1-13) (0.009)
12	wild-type	1. wild-type (97.781) 2. pre-S1 del (2.099) 3. pre-S2 del (0.112) 4. pre-S1+pre-S2 del (0.008) I. pre-S1 plus pre-S2 del (2.211) II. pre-S1 plus pre-S1+pre-S2 del (2.107) III. pre-S2 plus pre-S1+pre-S2 del (0.120) IV. all three types of pre-S del (2.219)	1. wild-type (97.781) 2. pre-S1 del (nt 3026-3205) (1.099) 3. pre-S2 del (nt 3211-3216) (0.055) 4. pre-S1+pre-S2 del (nt 3026-3205, 43-138) (0.002)
13	wild-type	1. wild-type (92.291) 2. pre-S1 del (4.092) 3. pre-S2 del (3.038) 4. pre-S1+pre-S2 del (0.578) I. pre-S1 plus pre-S2 del (7.130) II. pre-S1 plus pre-S1+pre-S2 del (4.670)	1. wild-type (92.291) 2. pre-S1 del (nt 2854-2970) (0.650) 3. pre-S2 del (nt 1-54) (2.641) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.435)

		III. pre-S2 plus pre-S1+pre-S2 del (3.616)	
		IV. all three types of pre-S del (7.709)	
14	wild-type	1. wild-type (69.001) 2. pre-S1 del (20.530) 3. pre-S2 del (9.463) 4. pre-S1+pre-S2 del (1.006) I. pre-S1 plus pre-S2 del (29.993) II. pre-S1 plus pre-S1+pre-S2 del (21.536) III. pre-S2 plus pre-S1+pre-S2 del (10.469) IV. all three types of pre-S del (31.999)	1. wild-type (69.001) 2. pre-S1 del (nt 3110-3127) (4.779) 3. pre-S2 del (nt 1-54) (8.226) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-18, 26-59) (0.259)
15	wild-type	1. wild-type (50.938) 2. pre-S1 del (19.760) 3. pre-S1+pre-S2 del (15.021) 4. pre-S2 del (14.280) I. pre-S1 plus pre-S2 del (34.040) II. pre-S1 plus pre-S1+pre-S2 del (34.781) III. pre-S2 plus pre-S1+pre-S2 del (29.301) IV. all three types of pre-S del (49.062)	1. wild-type (50.938) 2. pre-S1 del (nt 2854-2970) (5.470) 3. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (12.421) 4. pre-S2 del (nt 1-54) (13.606)
16	wild-type	1. wild-type (75.230) 2. pre-S1+pre-S2 del (13.878) 3. pre-S1 del (10.730) 4. pre-S2 del (0.162) I. pre-S1 plus pre-S2 del (10.892) II. pre-S1 plus pre-S1+pre-S2 del (24.608) III. pre-S2 plus pre-S1+pre-S2 del (14.040) IV. all three types of pre-S del (24.770)	1. wild-type (75.230) 2. pre-S1+pre-S2 del (nt 2860-2880, 2954-3097, 1-15) (11.572) 3. pre-S1 del (nt 2860-2880, 2954-3097) (9.550) 4. pre-S2 del (nt 1-15) (0.138)

17	wild-type	1. wild-type (98.571) 2. pre-S1 del (1.173) 3. pre-S2 del (0.178) 4. pre-S1+pre-S2 del (0.078) I. pre-S1 plus pre-S2 del (1.351) II. pre-S1 plus pre-S1+pre-S2 del (1.251) III. pre-S2 plus pre-S1+pre-S2 del (0.256) IV. all three types of pre-S del (1.429)	1. wild-type (98.571) 2. pre-S1 del (nt 3106-3129) (0.242) 3. pre-S2 del (nt 2-149) (0.043) 4. pre-S1+pre-S2 del (nt 2854-2985, 27-149) (0.014)
18	wild-type	1. wild-type (96.915) 2. pre-S1 del (2.701) 3. pre-S2 del (0.364) 4. pre-S1+pre-S2 del (0.020) I. pre-S1 plus pre-S2 del (3.065) II. pre-S1 plus pre-S1+pre-S2 del (2.721) III. pre-S2 plus pre-S1+pre-S2 del (0.384) IV. all three types of pre-S del (3.085)	1. wild-type (96.915) 2. pre-S1 del (nt 3026-3205) (1.755) 3. pre-S2 del (nt 25-54) (0.148) 4. pre-S1+pre-S2 del (nt 2854-2988, 45-149) (0.004)
19	wild-type	1. wild-type (98.069) 2. pre-S1 del (1.263) 3. pre-S2 del (0.609) 4. pre-S1+pre-S2 del (0.059) I. pre-S1 plus pre-S2 del (1.872) II. pre-S1 plus pre-S1+pre-S2 del (1.322) III. pre-S2 plus pre-S1+pre-S2 del (0.668) IV. all three types of pre-S del (1.931)	1. wild-type (98.069) 2. pre-S1 del (nt 3103-3126) (0.253) 3. pre-S2 del (nt 1-15) (0.535) 4. pre-S1+pre-S2 del (nt 2854-2996, 44-144) (0.015)

20	wild-type	1. wild-type (69.571) 2. pre-S2 del (18.590) 3. pre-S1+pre-S2 del (8.224) 4. pre-S1 del (3.615) I. pre-S1 plus pre-S2 del (22.205) II. pre-S1 plus pre-S1+pre-S2 del (11.839) III. pre-S2 plus pre-S1+pre-S2 del (26.814) IV. all three types of pre-S del (30.429)	1. wild-type (69.571) 2. pre-S2 del (nt 1-54) (18.482) 3. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (7.712) 4. pre-S1 del (nt 2855-2872) (2.674)
21	wild-type	1. pre-S2 del (41.477) 2. pre-S1+pre-S2 del (39.126) 3. wild-type (12.348) 4. pre-S1 del (7.048) I. pre-S1 plus pre-S2 del (48.525) II. pre-S1 plus pre-S1+pre-S2 del (46.174) III. pre-S2 plus pre-S1+pre-S2 del (80.603) IV. all three types of pre-S del (87.652)	1. pre-S2 del (nt 1-54) (24.620) 2. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (26.451) 3. wild-type (12.348) 4. pre-S1 del (nt 2855-2872) (4.978)
22	1. wild-type 2. pre-S1 del (nt 2855-2872)	1. pre-S2 del (42.909) 2. pre-S1+pre-S2 del (27.915) 3. wild-type (17.564) 4. pre-S1 del (11.612) I. pre-S1 plus pre-S2 del (54.521) II. pre-S1 plus pre-S1+pre-S2 del (39.527) III. pre-S2 plus pre-S1+pre-S2 del (70.824) IV. all three types of pre-S del (82.436)	1. pre-S2 del (nt 1-54) (38.091) 2. pre-S1+pre-S2 del (2855-2872, 1-54) (25.090) 3. wild-type (17.564) 4. pre-S1 del (nt 2855-2872) (8.072)

23	1. wild-type 2. pre-S1+pre-S2 del (nt 2855-2872, 1-54)	1. wild-type (52.661) 2. pre-S2 del (25.093) 3. pre-S1+pre-S2 del (11.424) 4. pre-S1 del (10.821) I. pre-S1 plus pre-S2 del (36.724) II. pre-S1 plus pre-S1+pre-S2 del (22.245) III. pre-S2 plus pre-S1+pre-S2 del (36.517) IV. all three types of pre-S del (47.339)	1. wild-type (52.661) 2. pre-S2 del (nt 1-54) (18.682) 3. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (8.344) 4. pre-S1 del (nt 2855-2872) (2.817)
24	wild-type	1. wild-type (96.073) 2. pre-S1 del (1.895) 3. pre-S1+pre-S2 del (1.202) 4. pre-S2 del (0.830) I. pre-S1 plus pre-S2 del (2.725) II. pre-S1 plus pre-S1+pre-S2 del (3.097) III. pre-S2 plus pre-S1+pre-S2 del (2.032) IV. all three types of pre-S del (3.927)	1. wild-type (96.073) 2. pre-S1 del (nt 2954-3097) (0.632) 3. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.452) 4. pre-S2 del (nt 1-54) (0.382)
25	wild-type	1. wild-type (93.547) 2. pre-S1 del (4.345) 3. pre-S2 del (1.844) 4. pre-S1+pre-S2 del (0.265) I. pre-S1 plus pre-S2 del (6.189) II. pre-S1 plus pre-S1+pre-S2 del (4.610) III. pre-S2 plus pre-S1+pre-S2 del (2.109) IV. all three types of pre-S del (6.453)	1. wild-type (93.547) 2. pre-S1 del (nt 3026-3205) (0.869) 3. pre-S2 del (nt 1-54) (1.107) 4. pre-S1+pre-S2 del (nt 2855-2872, 2897-2923, 1-54) (0.180)

26	wild-type	1. wild-type (97.921) 2. pre-S1 del (1.388) 3. pre-S2 del (0.691) 4. pre-S1+pre-S2 del (0.000) I. pre-S1 plus pre-S2 del (2.079) II. pre-S1 plus pre-S1+pre-S2 del (1.388) III. pre-S2 plus pre-S1+pre-S2 del (0.691) IV. all three types of pre-S del (2.079)	1. wild-type (97.921) 2. pre-S1 del (nt 3067-3162) (0.040) 3. pre-S2 del (nt 3211-3216) (0.046) 4. pre-S1+pre-S2 del (0.000)
27	pre-S1+pre-S2 del (nt 2855-2872, 1-54)	1. wild-type (60.195) 2. pre-S2 del (22.882) 3. pre-S1+pre-S2 del (9.280) 4. pre-S1 del (7.643) I. pre-S1 plus pre-S2 del (30.525) II. pre-S1 plus pre-S1+pre-S2 del (16.923) III. pre-S2 plus pre-S1+pre-S2 del (32.162) IV. all three types of pre-S del (39.805)	1. wild-type (60.195) 2. pre-S2 del (nt 1-57) (22.473) 3. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (4.815) 4. pre-S1 del (nt 3025-3126) (4.396)
28	wild-type	1. pre-S1 del (75.113) 2. wild-type (11.506) 3. pre-S1+pre-S2 del (10.472) 4. pre-S2 del (2.909) I. pre-S1 plus pre-S2 del (78.022) II. pre-S1 plus pre-S1+pre-S2 del (85.585) III. pre-S2 plus pre-S1+pre-S2 del (13.381) IV. all three types of pre-S del (88.494)	1. pre-S1 del (nt 2855-2872) (5.979) 2. wild-type (11.506) 3. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (8.016) 4. pre-S2 del (nt 1-54) (2.747)

29	wild-type	1. wild-type (97.730) 2. pre-S1 del (1.691) 3. pre-S2 del (0.575) 4. pre-S1+pre-S2 del (0.004) I. pre-S1 plus pre-S2 del (2.266) II. pre-S1 plus pre-S1+pre-S2 del (1.695) III. pre-S2 plus pre-S1+pre-S2 del (0.579) IV. all three types of pre-S del (2.270)	1. wild-type (97.730) 2. pre-S1 del (nt 2855-2970) (0.217) 3. pre-S2 del (nt 1-12) (0.323) 4. pre-S1+pre-S2 del (nt 2855-2970, 1-54) (0.004)
30	pre-S1 del (nt 3010-3075)	1. wild-type (87.018) 2. pre-S1 del (11.703) 3. pre-S2 del (1.113) 4. pre-S1+pre-S2 del (0.166) I. pre-S1 plus pre-S2 del (12.816) II. pre-S1 plus pre-S1+pre-S2 del (11.869) III. pre-S2 plus pre-S1+pre-S2 del (1.279) IV. all three types of pre-S del (12.982)	1. wild-type (87.018) 2. pre-S1 del (nt 3010-3075) (5.189) 3. pre-S2 del (nt 1-9) (0.392) 4. pre-S1+pre-S2 del (nt 3010-3075, 1-9) (0.046)
31	pre-S1 del (nt 2910-3089)	1. wild-type (78.709) 2. pre-S1 del (20.616) 3. pre-S2 del (0.646) 4. pre-S1+pre-S2 del (0.029) I. pre-S1 plus pre-S2 del (21.262) II. pre-S1 plus pre-S1+pre-S2 del (20.645) III. pre-S2 plus pre-S1+pre-S2 del (0.675) IV. all three types of pre-S del (21.291)	1. wild-type (78.709) 2. pre-S1 del (nt 2910-3089) (9.181) 3. pre-S2 del (nt 1-15) (0.345) 4. pre-S1+pre-S2 del (nt 2910-3055, 3067-3089, 4-12) (0.007)

32	wild-type	1. wild-type (98.105) 2. pre-S1 del (1.152) 3. pre-S2 del (0.733) 4. pre-S1+pre-S2 del (0.010) I. pre-S1 plus pre-S2 del (1.885) II. pre-S1 plus pre-S1+pre-S2 del (1.162) III. pre-S2 plus pre-S1+pre-S2 del (0.743) IV. all three types of pre-S del (1.895)	1. wild-type (98.105) 2. pre-S1 del (nt 2855-2970) (0.060) 3. pre-S2 del (nt 1-9) (0.408) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.003)
33	wild-type	1. wild-type (80.067) 2. pre-S2 del (9.477) 3. pre-S1 del (8.993) 4. pre-S1+pre-S2 del (1.463) I. pre-S1 plus pre-S2 del (18.470) II. pre-S1 plus pre-S1+pre-S2 del (10.456) III. pre-S2 plus pre-S1+pre-S2 del (10.940) IV. all three types of pre-S del (19.933)	1. wild-type (80.067) 2. pre-S2 del (nt 1-12) (8.957) 3. pre-S1 del (nt 2866-3075) (7.322) 4. pre-S1+pre-S2 del (nt 2866-3075, 1-9) (0.871)
34	wild-type	1. wild-type (80.175) 2. pre-S2 del (13.639) 3. pre-S1 del (5.169) 4. pre-S1+pre-S2 del (1.017) I. pre-S1 plus pre-S2 del (18.808) II. pre-S1 plus pre-S1+pre-S2 del (6.186) III. pre-S2 plus pre-S1+pre-S2 del (14.656) IV. all three types of pre-S del (19.825)	1. wild-type (80.175) 2. pre-S2 del (nt 1-30) (4.394) 3. pre-S1 del (nt 2865-2975) (2.356) 4. pre-S1+pre-S2 del (nt 2866-2975, 1-29) (0.212)

35	wild-type	1. wild-type (92.156) 2. pre-S1 del (7.022) 3. pre-S2 del (0.793) 4. pre-S1+pre-S2 del (0.029) I. pre-S1 plus pre-S2 del (7.815) II. pre-S1 plus pre-S1+pre-S2 del (7.051) III. pre-S2 plus pre-S1+pre-S2 del (0.822) IV. all three types of pre-S del (7.844)	1. wild-type (92.156) 2. pre-S1 del (nt 2854-3018) (1.346) 3. pre-S2 del (nt 1-9) (0.323) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-9) (0.004)
36	1. pre-S1 del (nt 2854-2970) 2. pre-S1+pre-S2 del (nt 2855-2872, 1-54)	1. pre-S1+pre-S2 del (40.433) 2. pre-S1 del (34.174) 3. wild-type (20.822) 4. pre-S2 del (4.571) I. pre-S1 plus pre-S2 del (38.745) II. pre-S1 plus pre-S1+pre-S2 del (74.607) III. pre-S2 plus pre-S1+pre-S2 del (45.004) IV. all three types of pre-S del (79.178)	1. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (22.713) 2. pre-S1 del (nt 2854-2970) (22.956) 3. wild-type (20.822) 4. pre-S2 del (nt 1-54) (4.380)
37	wild-type	1. wild-type (80.758) 2. pre-S2 del (12.910) 3. pre-S1 del (5.629) 4. pre-S1+pre-S2 del (0.704) I. pre-S1 plus pre-S2 del (18.539) II. pre-S1 plus pre-S1+pre-S2 del (6.333) III. pre-S2 plus pre-S1+pre-S2 del (13.614) IV. all three types of pre-S del (19.242)	1. wild-type (80.758) 2. pre-S2 del (nt 1-54) (6.944) 3. pre-S1 del (nt 2855-2875) (4.209) 4. pre-S1+pre-S2 del (nt 2855-2875, 1-54) (0.253)

38	pre-S1 del (nt 3021-3203)	1. wild-type (61.320) 2. pre-S1 del (34.045) 3. pre-S2 del (3.491) 4. pre-S1+pre-S2 del (1.144) I. pre-S1 plus pre-S2 del (37.536) II. pre-S1 plus pre-S1+pre-S2 del (35.189) III. pre-S2 plus pre-S1+pre-S2 del (4.635) IV. all three types of pre-S del (38.680)	1. wild-type (61.320) 2. pre-S1 del (nt 3021-3203) (23.818) 3. pre-S2 del (nt 6-41) (1.385) 4. pre-S1+pre-S2 del (nt 2855-2872, 2919-3126, 6-41) (0.163)
39	wild-type	1. wild-type (97.172) 2. pre-S1 del (1.957) 3. pre-S2 del (0.850) 4. pre-S1+pre-S2 del (0.021) I. pre-S1 plus pre-S2 del (2.807) II. pre-S1 plus pre-S1+pre-S2 del (1.978) III. pre-S2 plus pre-S1+pre-S2 del (0.871) IV. all three types of pre-S del (2.828)	1. wild-type (97.172) 2. pre-S1 del (nt 3021-3203) (0.353) 3. pre-S2 del (nt 1-12) (0.488) 4. pre-S1+pre-S2 del (nt 3078-3094, 6-41) (0.003)
40	wild-type	1. wild-type (97.501) 2. pre-S1 del (1.586) 3. pre-S2 del (0.908) 4. pre-S1+pre-S2 del (0.005) I. pre-S1 plus pre-S2 del (2.494) II. pre-S1 plus pre-S1+pre-S2 del (1.591) III. pre-S2 plus pre-S1+pre-S2 del (0.913) IV. all three types of pre-S del (2.499)	1. wild-type (97.501) 2. pre-S1 del (nt 3107-3202) (0.081) 3. pre-S2 del (nt 1-12) (0.529) 4. pre-S1+pre-S2 del (nt 3133-3136, 1-10) (0.005)

41	pre-S1 del (nt 2858-2986)	1. pre-S1 del (84.262) 2. wild-type (14.798) 3. pre-S1+pre-S2 del (0.779) 4. pre-S2 del (0.161) I. pre-S1 plus pre-S2 del (84.423) II. pre-S1 plus pre-S1+pre-S2 del (85.041) III. pre-S2 plus pre-S1+pre-S2 del (0.940) IV. all three types of pre-S del (85.202)	1. pre-S1 del (nt 2858-2986) (82.700) 2. wild-type (14.798) 3. pre-S1+pre-S2 del (nt 2858-2981, 1-10) (0.268) 4. pre-S2 del (nt 1-33) (0.053)
42	wild-type	1. wild-type (97.757) 2. pre-S1 del (1.305) 3. pre-S2 del (0.923) 4. pre-S1+pre-S2 del (0.015) I. pre-S1 plus pre-S2 del (2.228) II. pre-S1 plus pre-S1+pre-S2 del (1.320) III. pre-S2 plus pre-S1+pre-S2 del (0.938) IV. all three types of pre-S del (2.243)	1. wild-type (97.757) 2. pre-S1 del (nt 3106-3207) (0.060) 3. pre-S2 del (nt 3211-3213) (0.124) 4. pre-S1+pre-S2 del (nt 3069-3118, 3211-3212) (0.004)
43	wild-type	1. pre-S2 del (49.695) 2. wild-type (49.248) 3. pre-S1+pre-S2 del (0.814) 4. pre-S1 del (0.242) I. pre-S1 plus pre-S2 del (49.937) II. pre-S1 plus pre-S1+pre-S2 del (1.056) III. pre-S2 plus pre-S1+pre-S2 del (50.509) IV. all three types of pre-S del (50.752)	1. pre-S2 del (nt 37-54) (43.929) 2. wild-type (49.248) 3. pre-S1+pre-S2 del (nt 3139-3142, 37-54) (0.049) 4. pre-S1 del (nt 3103-3200) (0.019)

44	wild-type	1. wild-type (98.244) 2. pre-S1 del (1.113) 3. pre-S2 del (0.643) 4. pre-S1+pre-S2 del (0.000) I. pre-S1 plus pre-S2 del (1.756) II. pre-S1 plus pre-S1+pre-S2 del (1.113) III. pre-S2 plus pre-S1+pre-S2 del (0.643) IV. all three types of pre-S del (1.756)	1. wild-type (98.244) 2. pre-S1 del (nt 3089-3202) (0.049) 3. pre-S2 del (nt 1-15) (0.319) 4. pre-S1+pre-S2 del (0.000)
45	wild-type	1. wild-type (97.534) 2. pre-S1 del (1.376) 3. pre-S2 del (1.076) 4. pre-S1+pre-S2 del (0.014) I. pre-S1 plus pre-S2 del (2.452) II. pre-S1 plus pre-S1+pre-S2 del (1.390) III. pre-S2 plus pre-S1+pre-S2 del (1.090) IV. all three types of pre-S del (2.466)	1. wild-type (97.534) 2. pre-S1 del (nt 3106-3207) (0.063) 3. pre-S2 del (nt 1-9) (0.769) 4. pre-S1+pre-S2 del (nt 3095-3150, 1-12) (0.007)
46	wild-type	1. wild-type (97.735) 2. pre-S1 del (1.534) 3. pre-S2 del (0.731) 4. pre-S1+pre-S2 del (0.000) I. pre-S1 plus pre-S2 del (2.265) II. pre-S1 plus pre-S1+pre-S2 del (1.534) III. pre-S2 plus pre-S1+pre-S2 del (0.731) IV. all three types of pre-S del (2.265)	1. wild-type (97.735) 2. pre-S1 del (nt 3138-3197) (0.111) 3. pre-S2 del (nt 1-9) (0.371) 4. pre-S1+pre-S2 del (0.000)

47	wild-type	1. wild-type (78.334) 2. pre-S1 del (20.695) 3. pre-S2 del (0.866) 4. pre-S1+pre-S2 del (0.105) I. pre-S1 plus pre-S2 del (21.561) II. pre-S1 plus pre-S1+pre-S2 del (20.800) III. pre-S2 plus pre-S1+pre-S2 del (0.971) IV. all three types of pre-S del (21.666)	1. wild-type (78.334) 2. pre-S1 del (nt 2895-3140) (15.887) 3. pre-S2 del (nt 1-9) (0.279) 4. pre-S1+pre-S2 del (nt 2895-3141, 1-19, 23-27) (0.023)
48	1. pre-S1 del (nt 2968-3093) 2. pre-S2 del (nt 1-54)	1. pre-S1 del (57.159) 2. wild-type (20.456) 3. pre-S2 del (13.666) 4. pre-S1+pre-S2 del (8.719) I. pre-S1 plus pre-S2 del (70.825) II. pre-S1 plus pre-S1+pre-S2 del (65.878) III. pre-S2 plus pre-S1+pre-S2 del (22.385) IV. all three types of pre-S del (79.544)	1. pre-S1 del (nt 2968-3093) (47.067) 2. wild-type (20.456) 3. pre-S2 del (nt 1-54) (8.120) 4. pre-S1+pre-S2 del (nt 2968-3093, 1-54) (3.469)
49	wild-type	1. wild-type (75.021) 2. pre-S2 del (21.983) 3. pre-S1 del (2.327) 4. pre-S1+pre-S2 del (0.669) I. pre-S1 plus pre-S2 del (24.310) II. pre-S1 plus pre-S1+pre-S2 del (2.996) III. pre-S2 plus pre-S1+pre-S2 del (22.652) IV. all three types of pre-S del (24.979)	1. wild-type (75.021) 2. pre-S2 del (nt 1-1) (19.454) 3. pre-S1 del (nt 2855-2872) (0.704) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.606)

50	wild-type	1. wild-type (98.169) 2. pre-S1 del (1.138) 3. pre-S2 del (0.693) 4. pre-S1+pre-S2 del (0.000) I. pre-S1 plus pre-S2 del (1.831) II. pre-S1 plus pre-S1+pre-S2 del (1.138) III. pre-S2 plus pre-S1+pre-S2 del (0.693) IV. all three types of pre-S del (1.831)	1. wild-type (98.169) 2. pre-S1 del (nt 3138-3197) (0.067) 3. pre-S2 del (nt 1-9) (0.377) 4. pre-S1+pre-S2 del (0.000)
51	wild-type	1. wild-type (97.734) 2. pre-S1 del (1.546) 3. pre-S2 del (0.719) 4. pre-S1+pre-S2 del (0.000) I. pre-S1 plus pre-S2 del (2.265) II. pre-S1 plus pre-S1+pre-S2 del (1.546) III. pre-S2 plus pre-S1+pre-S2 del (0.719) IV. all three types of pre-S del (2.266)	1. wild-type (97.734) 2. pre-S1 del (nt 3104-3202) (0.165) 3. pre-S2 del (nt 1-9) (0.377) 4. pre-S1+pre-S2 del (0.000)
52	wild-type	1. wild-type (98.167) 2. pre-S1 del (1.192) 3. pre-S2 del (0.617) 4. pre-S1+pre-S2 del (0.024) I. pre-S1 plus pre-S2 del (1.809) II. pre-S1 plus pre-S1+pre-S2 del (1.216) III. pre-S2 plus pre-S1+pre-S2 del (0.641) IV. all three types of pre-S del (1.833)	1. wild-type (98.167) 2. pre-S1 del (nt 2854-3128) (0.132) 3. pre-S2 del (nt 1-9) (0.246) 4. pre-S1+pre-S2 del (nt 3022-3125, 1-55) (0.018)

53	wild-type	1. wild-type (97.002) 2. pre-S1 del (2.144) 3. pre-S2 del (0.849) 4. pre-S1+pre-S2 del (0.005) I. pre-S1 plus pre-S2 del (2.993) II. pre-S1 plus pre-S1+pre-S2 del (2.149) III. pre-S2 plus pre-S1+pre-S2 del (0.854) IV. all three types of pre-S del (2.998)	1. wild-type (97.002) 2. pre-S1 del (nt 3026-3205) (0.927) 3. pre-S2 del (nt 1-9) (0.528) 4. pre-S1+pre-S2 del (nt 3020-3119, 1-55) (0.005)
54	wild-type	1. wild-type (96.687) 2. pre-S1 del (2.625) 3. pre-S2 del (0.676) 4. pre-S1+pre-S2 del (0.013) I. pre-S1 plus pre-S2 del (3.301) II. pre-S1 plus pre-S1+pre-S2 del (2.638) III. pre-S2 plus pre-S1+pre-S2 del (0.689) IV. all three types of pre-S del (3.313)	1. wild-type (96.687) 2. pre-S1 del (nt 2855-2872) (1.117) 3. pre-S2 del (nt 1-12) (0.411) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-17) (0.004)
55	wild-type	1. wild-type (64.666) 2. pre-S1 del (34.582) 3. pre-S2 del (0.493) 4. pre-S1+pre-S2 del (0.259) I. pre-S1 plus pre-S2 del (35.075) II. pre-S1 plus pre-S1+pre-S2 del (34.841) III. pre-S2 plus pre-S1+pre-S2 del (0.752) IV. all three types of pre-S del (35.334)	1. wild-type (64.666) 2. pre-S1 del (nt 2858-2986) (33.466) 3. pre-S2 del (nt 1-9) (0.303) 4. pre-S1+pre-S2 del (nt 2858-2981, 1-10) (0.069)

56	wild-type	1. pre-S1 del (69.372) 2. wild-type (30.404) 3. pre-S2 del (0.218) 4. pre-S1+pre-S2 del (0.005) I. pre-S1 plus pre-S2 del (69.590) II. pre-S1 plus pre-S1+pre-S2 del (69.377) III. pre-S2 plus pre-S1+pre-S2 del (0.223) IV. all three types of pre-S del (69.596)	1. pre-S1 del (nt 2856-3101) (66.854) 2. wild-type (30.404) 3. pre-S2 del (nt 1-12) (0.130) 4. pre-S1+pre-S2 del (nt 2984-3098, 1-10) (0.002)
57	wild-type	1. wild-type (94.005) 2. pre-S1 del (3.729) 3. pre-S2 del (1.824) 4. pre-S1+pre-S2 del (0.442) I. pre-S1 plus pre-S2 del (5.553) II. pre-S1 plus pre-S1+pre-S2 del (4.171) III. pre-S2 plus pre-S1+pre-S2 del (2.266) IV. all three types of pre-S del (5.995)	1. wild-type (94.005) 2. pre-S1 del (nt 2895-3188) (0.677) 3. pre-S2 del (nt 1-54) (1.029) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.253)
58	wild-type	1. wild-type (95.575) 2. pre-S1 del (2.481) 3. pre-S2 del (1.931) 4. pre-S1+pre-S2 del (0.013) I. pre-S1 plus pre-S2 del (4.412) II. pre-S1 plus pre-S1+pre-S2 del (2.494) III. pre-S2 plus pre-S1+pre-S2 del (1.944) IV. all three types of pre-S del (4.424)	1. wild-type (95.575) 2. pre-S1 del (nt 3039-3092) (0.590) 3. pre-S2 del (nt 1-12) (0.510) 4. pre-S1+pre-S2 del (nt 2854-2940, 1-144) (0.007)

59	pre-S2 del (nt 15-56)	1. pre-S2 del (64.182) 2. wild-type (32.367) 3. pre-S1+pre-S2 del (2.975) 4. pre-S1 del (0.476) I. pre-S1 plus pre-S2 del (64.658) II. pre-S1 plus pre-S1+pre-S2 del (3.451) III. pre-S2 plus pre-S1+pre-S2 del (67.157) IV. all three types of pre-S del (67.633)	1. pre-S2 del (nt 15-56) (33.613) 2. wild-type (32.367) 3. pre-S1+pre-S2 del (nt 3088-3126, 15-56) (0.176) 4. pre-S1 del (nt 2923-3090) (0.073)
60	wild-type	1. pre-S1 del (38.192) 2. wild-type (34.644) 3. pre-S2 del (19.195) 4. pre-S1+pre-S2 del (7.968) I. pre-S1 plus pre-S2 del (57.387) II. pre-S1 plus pre-S1+pre-S2 del (46.160) III. pre-S2 plus pre-S1+pre-S2 del (27.163) IV. all three types of pre-S del (65.356)	1. pre-S1 del (nt 2854-2970) (27.998) 2. wild-type (30.404) 3. pre-S2 del (nt 1-54) (18.373) 4. pre-S1+pre-S2 del (nt 3025-3126, 1-57) (2.383)
61	pre-S1 del (nt 2855-2965)	1. pre-S1 del (40.086) 2. wild-type (34.919) 3. pre-S2 del (16.065) 4. pre-S1+pre-S2 del (8.930) I. pre-S1 plus pre-S2 del (56.151) II. pre-S1 plus pre-S1+pre-S2 del (49.016) III. pre-S2 plus pre-S1+pre-S2 del (24.995) IV. all three types of pre-S del (65.081)	1. pre-S1 del (nt 2855-2965) (24.125) 2. wild-type (34.919) 3. pre-S2 del (nt 1-54) (15.565) 4. pre-S1+pre-S2 del (nt 3022-3126, 1-60) (2.911)

62	pre-S1+pre-S2 del (nt 2855-2872, 1-54)	1. pre-S1 del (29.181) 2. wild-type (23.836) 3. pre-S2 del (23.645) 4. pre-S1+pre-S2 del (23.338) I. pre-S1 plus pre-S2 del (52.826) II. pre-S1 plus pre-S1+pre-S2 del (52.519) III. pre-S2 plus pre-S1+pre-S2 del (46.983) IV. all three types of pre-S del (76.164)	1. pre-S1 del (nt 2856-2969) (17.015) 2. wild-type (23.836) 3. pre-S2 del (nt 1-54) (23.402) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (15.956)
63	pre-S1 del (nt 2854-2970)	1. pre-S1 del (52.401) 2. wild-type (42.979) 3. pre-S2 del (2.312) 4. pre-S1+pre-S2 del (2.308) I. pre-S1 plus pre-S2 del (54.713) II. pre-S1 plus pre-S1+pre-S2 del (54.709) III. pre-S2 plus pre-S1+pre-S2 del (4.620) IV. all three types of pre-S del (57.021)	1. pre-S1 del (nt 2854-2970) (43.037) 2. wild-type (42.979) 3. pre-S2 del (nt 1-54) (1.977) 4. pre-S1+pre-S2 del (nt 2860-2880, 2954-3097, 1-15) (1.445)
64	wild-type	1. pre-S1 del (41.155) 2. wild-type (37.573) 3. pre-S2 del (18.905) 4. pre-S1+pre-S2 del (2.367) I. pre-S1 plus pre-S2 del (60.060) II. pre-S1 plus pre-S1+pre-S2 del (43.522) III. pre-S2 plus pre-S1+pre-S2 del (21.272) IV. all three types of pre-S del (62.427)	1. pre-S1 del (nt 2854-2970) (35.773) 2. wild-type (37.573) 3. pre-S2 del (nt 1-54) (18.494) 4. pre-S1+pre-S2 del (nt 2855-2970, 1-54) (1.676)

65	wild-type	1. pre-S1 del (63.334) 2. wild-type (32.806) 3. pre-S2 del (1.965) 4. pre-S1+pre-S2 del (1.895) I. pre-S1 plus pre-S2 del (65.229) II. pre-S1 plus pre-S1+pre-S2 del (65.229) III. pre-S2 plus pre-S1+pre-S2 del (3.860) IV. all three types of pre-S del (67.194)	1. pre-S1 del (nt 2854-2970) (33.450) 2. wild-type (32.806) 3. pre-S2 del (nt 1-54) (1.708) 4. pre-S1+pre-S2 del (nt 2855-2970, 1-54) (0.994)
66	pre-S1+pre-S2 del (nt 2855-2872, 1-54)	1. wild-type (30.973) 2. pre-S1+pre-S2 del (27.774) 3. pre-S1 del (27.161) 4. pre-S2 del (14.091) I. pre-S1 plus pre-S2 del (41.252) II. pre-S1 plus pre-S1+pre-S2 del (54.935) III. pre-S2 plus pre-S1+pre-S2 del (41.865) IV. all three types of pre-S del (69.027)	1. wild-type (30.973) 2. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (24.398) 3. pre-S1 del (nt 2854-2970) (14.813) 4. pre-S2 del (nt 1-54) (13.846)
67	wild-type	1. wild-type (46.914) 2. pre-S1 del (45.517) 3. pre-S2 del (6.834) 4. pre-S1+pre-S2 del (0.735) I. pre-S1 plus pre-S2 del (52.351) II. pre-S1 plus pre-S1+pre-S2 del (46.252) III. pre-S2 plus pre-S1+pre-S2 del (7.569) IV. all three types of pre-S del (53.086)	1. wild-type (46.914) 2. pre-S1 del (nt 2854-2970) (37.666) 3. pre-S2 del (nt 1-54) (5.216) 4. pre-S1+pre-S2 del (nt 2854-2970, 1-54) (0.307)

68	pre-S1+pre-S2 del (nt 2855-2872, 1-54)	1. pre-S1 del (43.130) 2. wild-type (41.965) 3. pre-S2 del (9.508) 4. pre-S1+pre-S2 del (5.397) I. pre-S1 plus pre-S2 del (52.638) II. pre-S1 plus pre-S1+pre-S2 del (48.527) III. pre-S2 plus pre-S1+pre-S2 del (14.905) IV. all three types of pre-S del (58.035)	1. pre-S1 del (nt 2854-2970) (35.888) 2. wild-type (41.965) 3. pre-S2 del (nt 1-54) (9.239) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (4.160)
69	pre-S1+pre-S2 del (nt 2855-2872, 1-54)	1. wild-type (36.868) 2. pre-S1 del (35.238) 3. pre-S2 del (21.130) 4. pre-S1+pre-S2 del (6.763) I. pre-S1 plus pre-S2 del (56.368) II. pre-S1 plus pre-S1+pre-S2 del (42.001) III. pre-S2 plus pre-S1+pre-S2 del (27.893) IV. all three types of pre-S del (63.132)	1. wild-type (36.868) 2. pre-S1 del (nt 2854-2970) (29.499) 3. pre-S2 del (nt 1-54) (20.218) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (3.332)
70	pre-S1 del (nt 2854-3021)	1. wild-type (94.788) 2. pre-S1 del (4.643) 3. pre-S2 del (0.455) 4. pre-S1+pre-S2 del (0.114) I. pre-S1 plus pre-S2 del (5.098) II. pre-S1 plus pre-S1+pre-S2 del (4.757) III. pre-S2 plus pre-S1+pre-S2 del (0.569) IV. all three types of pre-S del (5.212)	1. wild-type (94.788) 2. pre-S1 del (nt 2854-3021) (3.404) 3. pre-S2 del (nt 1-15) (0.192) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.086)

71	wild-type	1. pre-S2 del (54.981) 2. wild-type (42.238) 3. pre-S1+pre-S2 del (2.274) 4. pre-S1 del (0.507) I. pre-S1 plus pre-S2 del (55.488) II. pre-S1 plus pre-S1+pre-S2 del (2.781) III. pre-S2 plus pre-S1+pre-S2 del (57.255) IV. all three types of pre-S del (57.762)	1. pre-S2 del (nt 48-56) (52.982) 2. wild-type (42.238) 3. pre-S1+pre-S2 del (nt 3026-3205, 48-56) (0.916) 4. pre-S1 del (nt 2854-3021) (0.334)
72	pre-S1 del (nt 2854-3021)	1. wild-type (91.511) 2. pre-S1 del (7.310) 3. pre-S2 del (0.807) 4. pre-S1+pre-S2 del (0.373) I. pre-S1 plus pre-S2 del (8.117) II. pre-S1 plus pre-S1+pre-S2 del (7.683) III. pre-S2 plus pre-S1+pre-S2 del (1.180) IV. all three types of pre-S del (8.489)	1. wild-type (91.511) 2. pre-S1 del (nt 2854-3021) (5.744) 3. pre-S2 del (nt 48-56) (0.294) 4. pre-S1+pre-S2 del (nt 2855-2872, 1-54) (0.322)
73	pre-S1+pre-S2 del (nt 2855-2872, 3012-3086, 1-51)	1. wild-type (52.297) 2. pre-S1+pre-S2 del (31.339) 3. pre-S1 del (12.863) 4. pre-S2 del (3.502) I. pre-S1 plus pre-S2 del (16.365) II. pre-S1 plus pre-S1+pre-S2 del (44.202) III. pre-S2 plus pre-S1+pre-S2 del (34.841) IV. all three types of pre-S del (47.703)	1. wild-type (52.297) 2. pre-S1+pre-S2 del (nt 2855-2872, 3012-3086, 1-51) (8.935) 3. pre-S1 del (nt 2855-2872, 3012-3092) (6.480) 4. pre-S2 del (nt 1-51) (2.094)

74	pre-S2 del (nt 1-57)	1. wild-type (58.984) 2. pre-S2 del (34.533) 3. pre-S1+pre-S2 del (4.497) 4. pre-S1 del (1.986) I. pre-S1 plus pre-S2 del (36.519) II. pre-S1 plus pre-S1+pre-S2 del (6.483) III. pre-S2 plus pre-S1+pre-S2 del (39.030) IV. all three types of pre-S del (41.016)	1. wild-type (58.984) 2. pre-S2 del (nt 1-57) (20.926) 3. pre-S1+pre-S2 del (nt 3026-3205, 2-55) (0.590) 4. pre-S1 del (nt 2944-3120) (1.128)
75	wild-type	1. wild-type (97.112) 2. pre-S1 del (2.178) 3. pre-S2 del (0.679) 4. pre-S1+pre-S2 del (0.031) I. pre-S1 plus pre-S2 del (2.857) II. pre-S1 plus pre-S1+pre-S2 del (2.209) III. pre-S2 plus pre-S1+pre-S2 del (0.710) IV. all three types of pre-S del (2.888)	1. wild-type (97.112) 2. pre-S1 del (nt 2854-3021) (0.641) 3. pre-S2 del (nt 25-54) (0.355) 4. pre-S1+pre-S2 del (nt 2855-2970, 25-54) (0.007)

^aAll the PCR bands visualized in agarose gel were analyzed and listed in descending size order.

^bThe total frequency of pre-S gene DNA in each type of pre-S deletion was shown in descending order (1-4).

^cThe pre-S gene DNA with the highest frequency in each type of pre-S deletion was shown.

^dThe pre-S deletion type above the cut-off percentage (4.643) was shown in bold.

^eCombination of different types of pre-S deletions was shown (I-IV).

Abbreviations: nt, nucleotide; del, deletion.

Table S2. Univariate and multivariate analyses of pre-S deletion type for overall survival in 75 HBV-related HCC patients

Characteristics	Univariate Analysis			Multivariate Analysis		
	HR	95% CI	P value	HR	95% CI	P value
Age (years) (>50 vs. ≤50)	0.764	0.270-2.159	0.6109			
Gender (men vs. women)	1.462	0.191-11.179	0.7143			
Smoking (yes vs. no)	1.004	0.357-2.821	0.9947			
Alcohol (yes vs. no)	2.048	0.760-5.524	0.1566			
HBsAg (positive vs. negative) ^a						
HBeAg (positive vs. negative) ^b	0.635	0.083-4.861	0.6616			
HBV genotype (B vs. C)	1.296	0.292-5.752	0.7332			
HBV DNA (IU/mL) (>1×10 ⁴ vs. ≤1×10 ⁴) ^c	1.682	0.561-5.043	0.3537			
Albumin (g/dL) (>3.8 vs. ≤3.8)	0.425	0.135-1.341	0.1444			
AST (U/L) (>34 vs. ≤34)	1.323	0.298-5.864	0.7126			
ALT (U/L) (>40 vs. ≤40)	0.849	0.307-2.349	0.7530			
AFP (ng/mL) (>400 vs. ≤400)	2.542	0.937-6.895	0.0669			
Tumor size (cm) (>5 vs. ≤5)	1.605	0.581-4.437	0.3615			
Tumor encapsulation (yes vs. no) ^d	0.386	0.124-1.199	0.0997			
Lymph node involvement (yes vs. no)	0.513	0.067-3.899	0.5186			
Portal vein thrombosis (yes vs. no)	1.981	0.444-8.834	0.3701			
Vascular invasion (yes vs. no)	1.930	0.680-5.480	0.2169			
Distant metastasis (yes vs. no)	3.336	0.924-12.040	0.0658			
Steatosis grade (2/3 vs. 0/1) ^e	21.354	1.329-343.018	0.0307*			
Metavir inflammation score (2/3 vs. 0/1) ^f	1.474	0.284-7.645	0.6438			
Ishak fibrosis score (4/5/6 vs. 0/1/2/3) ^g	1.153	0.324-4.100	0.8263			
Child-Pugh cirrhosis score (B/C vs. A)	2.953	1.044-8.356	0.0413*	2.974	1.035-8.549	0.0430*
CLIP score (4/5/6 vs. 0/1/2/3)	5.233	0.675-40.579	0.1133			

Tumor differentiation grade (3/4 vs. 1/2)	0.818	0.291-2.301	0.7040			
BCLC stage (C/D vs. A/B)	1.314	0.296-5.834	0.7199			
AJCC TNM stage (IIIA/IIIB/IIIC/IVA/IVB vs. I/II)	3.113	1.053-9.201	0.0400*	3.146	1.041-9.510	0.0423*
Antiviral therapy after surgery (yes vs. no)	0.746	0.279-1.993	0.5590			
Del spanning pre-S gene segments (yes vs. no)	1.051	0.373-2.962	0.9250			
Only pre-S1 del (yes vs. no)	1.324	0.454-3.868	0.6073			
Only pre-S2 del (yes vs. no)	0.814	0.107-6.206	0.8423			
Only pre-S1+pre-S2 del (yes vs. no) ^h						
Both pre-S1 and pre-S2 del (yes vs. no)	3.216	0.708-14.615	0.1304			
Both pre-S1 and pre-S1+pre-S2 del (yes vs. no)	1.606	0.209-12.326	0.6487			
Both pre-S2 and pre-S1+pre-S2 del (yes vs. no)	0.000	0.000	0.9937			
All three types of pre-S del (yes vs. no)	0.302	0.039-2.366	0.2542			
Del spanning the pre-S1 gene segment (yes vs. no)	1.110	0.402-3.065	0.8407			
Del spanning the pre-S2 gene segment (yes vs. no)	0.818	0.283-2.366	0.7104			

^aThere were no patients negative for HBsAg for analysis.

^bOnly 71 patients with available data were analyzed.

^cOnly 74 patients with available data were analyzed.

^dOnly 62 patients with available data were analyzed.

^eOnly 25 patients with available data were analyzed and thus excluded from multivariate analysis.

^fOnly 44 patients with available data were analyzed.

^gOnly 56 patients with available data were analyzed.

^hThere were no patients with only pre-S1+pre-S2 del for analysis.

*, P value<0.05.

Abbreviations: HR, hazard ratio; CI, confidence interval; del, deletion.

Table S3. Univariate and multivariate analyses of pre-S deletion percentage for overall survival in 75 HBV-related HCC patients

Characteristics	Univariate Analysis			Multivariate Analysis		
	HR	95% CI	P value	HR	95% CI	P value
Child-Pugh cirrhosis score (B/C vs. A)	2.953	1.044-8.356	0.0413*	2.974	1.035-8.549	0.0430*
AJCC TNM stage (IIIA/IIIB/IIIC/IVA/IVB vs. I/II)	3.113	1.053-9.201	0.0400*	3.146	1.041-9.510	0.0423*
Pre-S1 del percentage ^a						
II (>1.895) vs. I (≤1.895) ^b	1.630	0.402-6.603	0.4938			
III (>5.629) vs. I (≤1.895)	1.061	0.214-5.263	0.9426			
IV (>28.270) vs. I (≤1.895)	1.396	0.306-6.367	0.6664			
Pre-S2 del percentage						
II (>0.643) vs. I (≤0.643)	1.482	0.387-5.669	0.5658			
III (>1.824) vs. I (≤0.643)	1.276	0.317-5.129	0.7315			
IV (>13.666) vs. I (≤0.643)	1.100	0.241-5.011	0.9024			
Pre-S1+pre-S2 del percentage						
II (>0.015) vs. I (≤0.015)	1.126	0.311-4.070	0.8567			
III (>0.442) vs. I (≤0.015)	0.766	0.171-3.439	0.7282			
IV (>2.975) vs. I (≤0.015)	0.717	0.157-3.287	0.6687			
Pre-S1 plus pre-S2 del percentage						
II (>2.725) vs. I (≤2.725)	1.846	0.473-7.209	0.3776			
III (>18.539) vs. I (≤2.725)	0.295	0.031-2.846	0.2913			
IV (>49.938) vs. I (≤2.725)	1.696	0.399-7.206	0.4745			
Pre-S1 plus pre-S1+pre-S2 del percentage						
II (>2.043) vs. I (≤2.043)	0.728	0.178-2.986	0.6595			
III (>6.332) vs. I (≤2.043)	1.108	0.297-4.136	0.8782			
IV (>44.201) vs. I (≤2.043)	0.767	0.164-3.589	0.7360			
Pre-S2 plus pre-S1+pre-S2 del percentage						

II (>0.689) vs. I (≤0.689)	1.819	0.500-6.615	0.3639
III (>2.109) vs. I (≤0.689)	1.137	0.283-4.563	0.8564
IV (>24.995) vs. I (≤0.689)	0.771	0.139-4.292	0.7668

All three types of pre-S del percentage

II (>2.828) vs. I (≤2.828)	1.035	0.288-3.718	0.9580
III (>21.666) vs. I (≤2.828)	0.208	0.023-1.866	0.1606
IV (>58.035) vs. I (≤2.828)	1.499	0.391-5.747	0.5548

^aPatients were divided into quarters of the distribution of indicated type of pre-S del percentage from the lowest to the highest (designated I to IV, respectively).

^bThreshold percentage of indicated pre-S del type for each quarter was shown in parenthesis.

*, P value<0.05.

Abbreviations: HR, hazard ratio; CI, confidence interval; del, deletion.

Table S4. Univariate and multivariate analyses of combined pre-S deletion for overall survival in 75 HBV-related HCC patients

Characteristics	Univariate Analysis			Multivariate Analysis		
	HR	95% CI	P value	HR	95% CI	P value
Child-Pugh cirrhosis score (B/C vs. A)	2.953	1.044-8.356	0.0413*	2.974	1.035-8.549	0.0430*
AJCC TNM stage (IIIA/IIIB/IIIC/IVA/IVB vs. I/II)	3.113	1.053-9.201	0.0400*	3.146	1.041-9.510	0.0423*
Combined pre-S del ^a						
com-II ^b vs. com-I						
com-III vs. com-I	1.101	0.297-4.074	0.8859			
com-IV vs. com-I	0.597	0.132-2.702	0.5029			

^aPatients were divided into four groups based on pre-S del types and percentages: com-I, del spanning the pre-S2 gene segment (no) & pre-S2 plus pre-S1+pre-S2 del percentage (I/II/III); com-II, del spanning the pre-S2 gene segment (no) & pre-S2 plus pre-S1+pre-S2 del percentage (IV); com-III, del spanning the pre-S2 gene segment (yes) & pre-S2 plus pre-S1+pre-S2 del percentage (I/II/III); and com-IV, del spanning the pre-S2 gene segment (yes) & pre-S2 plus pre-S1+pre-S2 del percentage (IV).

^bThere were no patients falling into the com-II group for analysis.

*, P value<0.05.

Abbreviations: HR, hazard ratio; CI, confidence interval; del, deletion; com, combined.