

1 **Supplementary Results**

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3 **Validation of microarray expression value by quantitative RT-PCR**

4 In order to validate the lncRNA expression value measured by microarray, we performed
 5 quantitative RT-PCR (qRT-PCR) of the three lncRNAs of the signature in 25 randomly selected
 6 tumor samples. Comparison of the expression values measured by qRT-PCR and microarray
 7 showed high level of positive correlation. The Pearson correlation coefficient of
 8 ENST00000435885.1, XLOC_013014, and ENST00000547963.1 were 0.74 (p<0.0001), 0.46
 9 (p=0.0211) and 0.51 (p=0.0089), respectively (Supplementary Figure 2). This result indicates that
 10 the lncRNA expression level measured by microarray is reliable.

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12 **Correlation of expression level among lncRNAs in the four-lncRNA and three-lncRNA signatures**

13 As mentioned in the method, in order to select the final prognostic signature, we compared the
 14 performances of k-lncRNA signature in the training set for all k=1,2,...,9 and the signatures with
 15 the best accuracies for each k was defined as the k-lncRNA signature. Three four-lncRNA
 16 signatures had the same highest accuracy (84.7%) for k=4 in the training set, which was also the
 17 highest accuracy for all k=1,2,...,9. The three four-lncRNA signatures were:

18 *Signature4_1*: ENST00000435885.1, XLOC_013014, XLOC_010016, and ENST00000547963.1

19 *Signature4_2*: ENST00000435885.1, XLOC_013014, XLOC_011774, and ENST00000547963.1

20 *Signature4_3*: ENST00000435885.1, XLOC_010016, XLOC_011774, and ENST00000547963.1

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22 Only one three-lncRNA signature had the highest accuracy (81.9%) for k=3 in the training set,
 23 which was:

24 *Signature3*: ENST00000435885.1, XLOC_013014, and ENST00000547963.1

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26 Two lncRNAs (ENST00000435885.1, ENST00000547963.1) were included in all the four-lncRNA
 27 signatures and the three-lncRNA signature, and XLOC_013014 were included in all these
 28 signatures except *Signature4_3*.

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30 We explored the correlations between these signature related lncRNAs by calculating the
 31 pearson and spearman's correlation coefficient in the training/test/independent sets, and the
 32 results are shown below.

33 Training set (a: p value < 2.2e⁻¹⁶, others: p value > 0.05)

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Pearson's correlation coefficient	ENST00000435885.1	XLOC_013014	ENST00000547963.1	XLOC_010016	XLOC_011774
ENST00000435885.1		-0.03	0.09	-0.04	-0.06
XLOC_013014			-0.12	0.93^a	0.95^a
ENST00000547963.1				-0.06	-0.07
XLOC_010016					0.95^a
XLOC_011774					

Spearman's correlation coefficient	ENST00000435885.1	XLOC_013014	ENST00000547963.1	XLOC_010016	XLOC_011774
ENST00000435885.1		-0.06	0.08	-0.12	-0.11
XLOC_013014			-0.05	0.86^a	0.91^a
ENST00000547963.1				0.03	0.04
XLOC_010016					0.92^a
XLOC_011774					

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2 Test set (a: p value < 2.2e⁻¹⁶, others: p value > 0.05)

Pearson's correlation coefficient	ENST00000435885.1	XLOC_013014	ENST00000547963.1	XLOC_010016	XLOC_011774
ENST00000435885.1		-0.04	0.13	-0.03	-0.04
XLOC_013014			-0.09	0.91^a	0.97^a
ENST00000547963.1				-0.17	-0.08
XLOC_010016					0.95^a
XLOC_011774					

Spearman's correlation coefficient	ENST00000435885.1	XLOC_013014	ENST00000547963.1	XLOC_010016	XLOC_011774
ENST00000435885.1		0.00	0.13	0.01	0.00
XLOC_013014			-0.09	0.92^a	0.96^a
ENST00000547963.1				-0.16	-0.07
XLOC_010016					0.95^a
XLOC_011774					

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5 Independent set (a: p value < 2.2e⁻¹⁶, b: p value = 7.66e⁻³, c: p value = 3.22e⁻², others: p value > 0.05)

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Pearson's correlation coefficient	ENST00000435885.1	XLOC_013014	ENST00000547963.1	XLOC_010016	XLOC_011774
ENST00000435885.1		0.08	0.34^b	0.00	0.10
XLOC_013014			0.05	0.90^a	0.99^a
ENST00000547963.1				0.06	0.08
XLOC_010016					0.88^a
XLOC_011774					

Spearman's correlation coefficient	ENST00000435885.1	XLOC_013014	ENST00000547963.1	XLOC_010016	XLOC_011774
ENST00000435885.1		0.15	0.28^c	0.00	0.17
XLOC_013014			0.10	0.89^a	0.99^a
ENST00000547963.1				0.09	0.09
XLOC_010016					0.88^a
XLOC_011774					

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9 From the results, high level of positive correlation (pearson's and spearman's correlation
10 coefficients of all pairs > 0.86, and p value < 2.2e⁻¹⁶) among the expression levels of XLOC_013014,
11 XLOC_010016 and XLOC_011774 were observed. So, there is one redundant lncRNA in each of
12 the four-lncRNA signatures. But for the three-lncRNA signature, there is no redundant lncRNA.

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14 **Comparison of prognostic performance between three-lncRNA and four-lncRNA signatures**

15 According to our algorithm, the patients could be classified into high- or low-risk groups by the
16 signature. The following tables show the classification results of patients in
17 training/test/independent sets by these signatures.

18 In the table, "1" denotes the patient is classified as low-risk and "2" denotes high-risk.

Training set

	Signature4_1	Signature4_2	Signature4_3	Signature3			Signature4_1	Signature4_2	Signature4_3	Signature3
p1	1	1	1	2		p31	1	1	1	2
p2	1	1	1	1		p32	1	1	1	1
p3	1	1	1	1		p33	2	2	2	2
p4	1	1	1	1		p34	2	2	2	2
p5	1	1	1	1		p35	2	2	2	2
p6	1	1	1	1		p36	1	1	1	1
p7	1	1	1	1		p37	1	1	1	1
p8	1	1	1	1		p38	2	2	2	2
p9	1	1	1	1		p39	2	2	2	2
p10	1	1	1	1		p40	2	2	2	2
p11	2	2	2	2		p41	1	1	1	1
p12	1	1	1	1		p42	2	2	2	2
p13	1	1	1	1		p43	2	2	2	2
p14	1	1	1	2		p44	2	2	2	2
p15	1	1	1	1		p45	2	2	2	2
p16	1	1	1	1		p46	2	2	2	2
p17	1	1	1	1		p47	2	2	2	2
p18	1	1	1	1		p48	2	2	2	2
p19	1	1	1	1		p49	2	2	2	2
p20	2	2	2	2		p50	2	2	2	2
p21	1	1	1	1		p51	2	2	2	2
p22	1	1	1	1		p52	1	1	1	1
p23	1	1	1	1		p53	2	2	2	2
p24	1	1	1	1		p54	2	2	2	2
p25	2	2	2	2		p55	2	2	2	2
p26	2	2	2	2		p56	2	2	2	2
p27	2	2	2	2		p57	2	2	2	2
p28	2	2	2	2		p58	1	1	1	1
p29	2	2	2	2		p59	1	1	1	1
p30	2	2	2	2		p60	2	2	2	2

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Test set

	Signature4_1	Signature4_2	Signature4_3	Signature3			Signature4_1	Signature4_2	Signature4_3	Signature3
p1	1	1	1	1		p31	1	1	1	1
p2	2	2	2	2		p32	2	2	2	2
p3	1	1	1	1		p33	2	2	2	2
p4	1	1	1	1		p34	1	1	1	1
p5	1	1	1	1		p35	2	2	2	2
p6	1	1	1	1		p36	1	1	1	1
p7	1	1	1	1		p37	2	2	2	2
p8	1	1	1	1		p38	2	2	2	2
p9	2	2	2	2		p39	1	1	1	2
p10	1	1	1	2		p40	1	1	1	1
p11	1	1	1	1		p41	1	1	1	1
p12	2	2	2	2		p42	1	1	1	2
p13	2	2	2	2		p43	1	1	1	1
p14	2	2	2	2		p44	2	2	2	2
p15	2	2	2	2		p45	1	1	1	1
p16	2	2	2	2		p46	1	1	1	1
p17	1	1	1	1		p47	1	1	1	1
p18	2	2	2	2		p48	1	1	1	1
p19	1	1	1	1		p49	1	1	1	1
p20	2	2	2	2		p50	2	2	2	2
p21	1	1	1	1		p51	2	2	2	2
p22	1	1	1	1		p52	2	2	2	2
p23	2	2	1	2		p53	1	1	1	1
p24	2	2	2	2		p54	1	1	1	1
p25	1	1	1	1		p55	1	1	1	1
p26	2	2	2	1		p56	1	1	1	1
p27	2	2	2	2		p57	1	1	1	1
p28	1	1	1	1		p58	1	2	2	1
p29	2	2	2	2		p59	1	1	1	1
p30	1	1	1	1						

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Independent set

	Signature4_1	Signature4_2	Signature4_3	Signature3			Signature4_1	Signature4_2	Signature4_3	Signature3
p1	2	2	2	2		p31	2	2	2	2
p2	2	2	2	2		p32	2	2	2	2
p3	2	2	2	1		p33	2	2	2	2
p4	2	2	2	2		p34	2	2	2	1
p5	2	2	2	2		p35	2	1	2	2
p6	1	1	1	1		p36	1	1	1	1
p7	1	1	1	1		p37	1	1	1	1
p8	2	2	2	2		p38	1	1	1	1
p9	1	1	1	1		p39	2	2	2	1
p10	2	2	2	2		p40	2	2	2	2
p11	1	1	1	1		p41	2	2	2	2
p12	1	1	1	1		p42	1	1	1	1
p13	1	1	1	1		p43	2	2	2	2
p14	1	1	1	1		p44	1	1	1	1
p15	2	2	2	2		p45	2	2	2	2
p16	1	1	1	1		p46	2	2	2	2
p17	1	1	1	1		p47	2	2	2	2
p18	1	1	1	1		p48	2	2	2	2
p19	2	2	2	2		p49	2	1	2	2
p20	2	2	2	2		p50	2	2	2	2
p21	2	2	2	1		p51	2	2	2	2
p22	2	2	2	2		p52	2	2	2	2
p23	2	2	2	2		p53	2	2	2	2
p24	2	2	2	2		p54	2	2	2	2
p25	1	1	1	1		p55	1	1	1	1
p26	2	1	2	2		p56	2	2	2	2
p27	1	1	2	2		p57	1	1	1	1
p28	1	1	1	1		p58	2	2	2	2
p29	2	2	2	2		p59	2	2	2	2
p30	2	2	2	2		p60	2	2	2	2

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From the tables, we see that the survival prediction results (high- or low-risk) of patients for the three four-lncRNA signatures were much the same. For the 60 patients of the training set, classification results of the three signatures were exactly the same. There were 1 (*Signature4_2*) and 2 (*Signature4_3*) patients with different classification compared to *Signature4_1* for the 59 patients of the test set, and were 3 (*Signature4_2*) and 1 (*Signature4_3*) for the 60 patients of the independent set.

Because the classification results for the four-lncRNA signatures were much the same, we only summarized the comparison between *Signature4_1* and the 3-lncRNA signature. From the tables, there were 3 patients with different classification compared to *Signature4_1* for 60 patients of the training set. The number was 4 and 5 for 59 patients of the test set and 60 patients of the independent set, respectively. So the classification results were very similar between the three four-lncRNA signatures and the three-lncRNA signature.

Prognostic performance comparison between three-lncRNA signature and four-lncRNA signatures

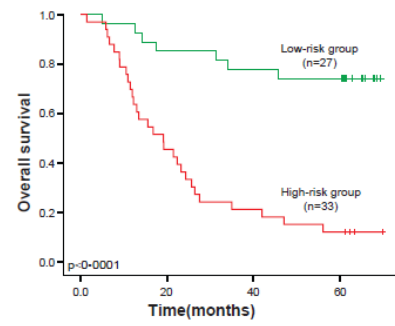
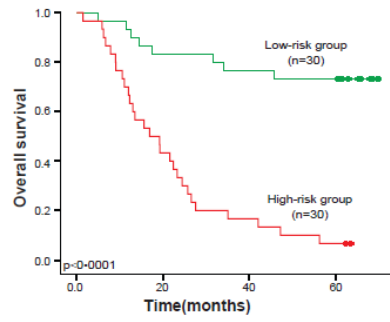
As mentioned above, the survival prediction results (high- or low-risk) of the three four-lncRNA signatures were very similar. So, we only show the performance comparison between *Signature4_1* and the three-lncRNA signature below. The prognostic performance of *Signature4_2/Signature4_3* and the three-lncRNA signature were also very similar (figure not shown).

Comparison of Kaplan-Meier survival curves of *Signature4_1* and *Signature3*

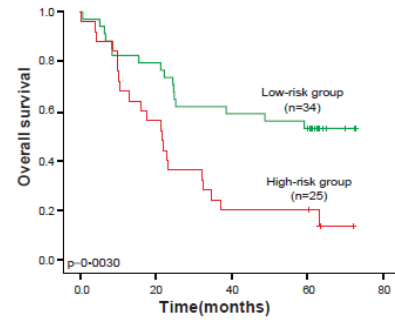
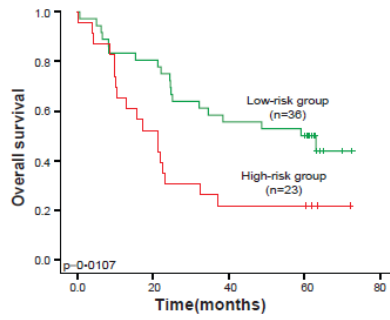
Signature4_1

Signature3

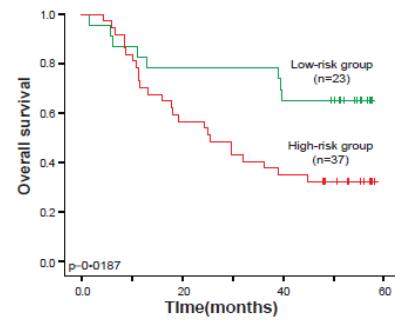
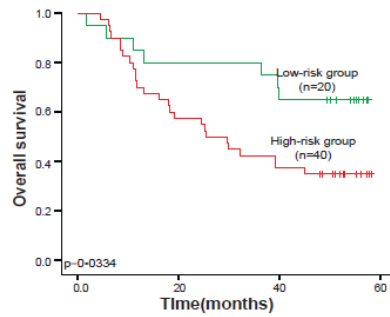
training set



test set



independent set



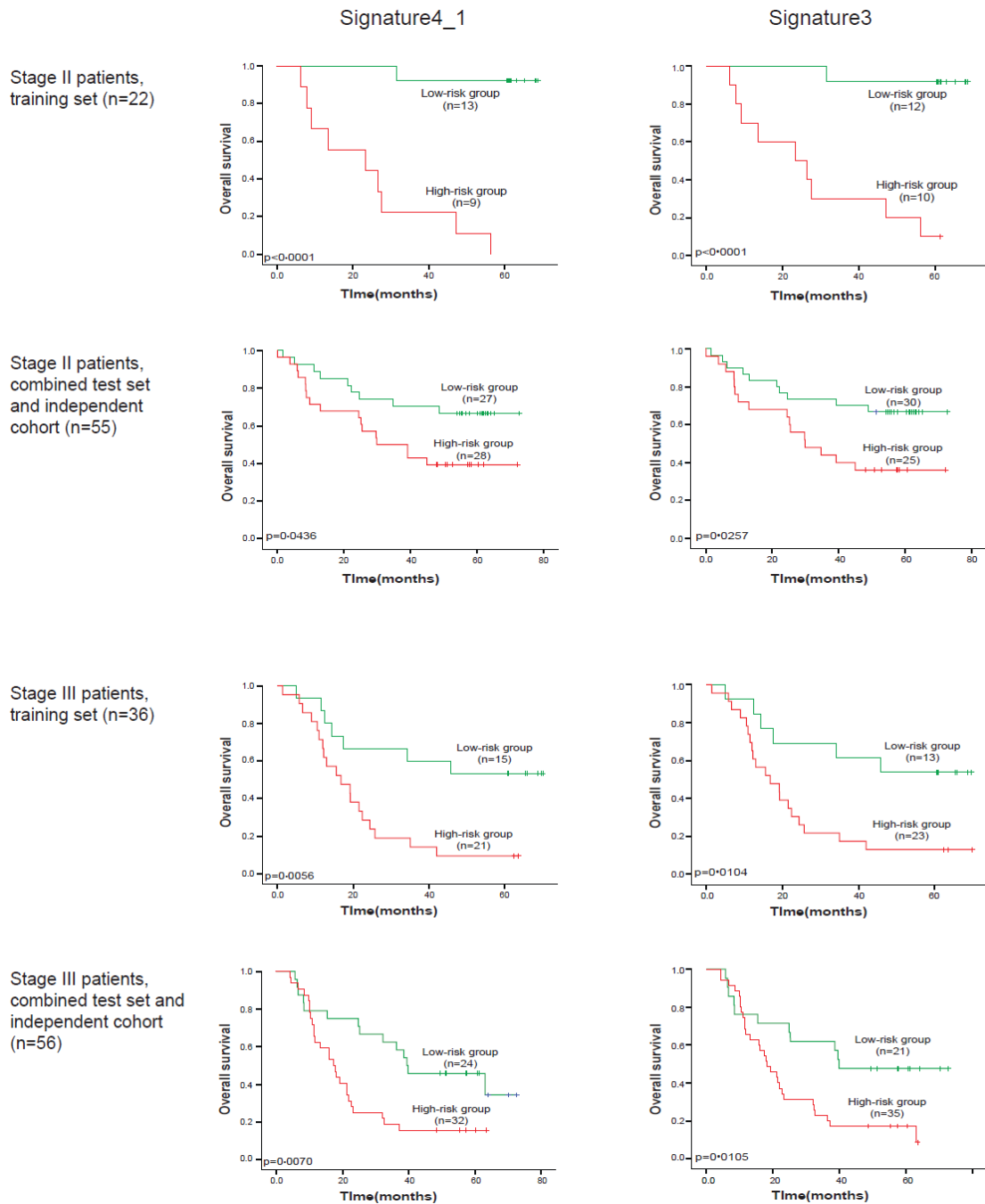
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Comparison of Kaplan-Meier survival curves of *Signature4_1* and *Signature3* within clinical stages



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3 From these results, we see that the performances between the four-lncRNA signature and
 4 three-lncRNA signature are very similar. The p values of Log-rank tests show that both the
 5 four-lncRNA signature and the three-lncRNA signature are able to predict the survival of patients
 6 both all together and within TNM stages and that four-lncRNA signature does not improve
 7 prognostic power compared with three-lncRNA signature.