

## SUPPLEMENTARY TABLES

**Supplementary Table 1. Correlation between ZFAS1 expression and clinicopathological features in the included colorectal cancer patients (*n* = 157).**

| Characteristics              | <i>n</i> = 157 | ZFAS1 expression         |                          | <i>P</i> -value | Adjusted OR(95%CI) |
|------------------------------|----------------|--------------------------|--------------------------|-----------------|--------------------|
|                              |                | Low (%)<br><i>n</i> =100 | High (%)<br><i>n</i> =57 |                 |                    |
| <b>Age</b>                   |                |                          |                          |                 |                    |
| <=64                         | 77             | 46(59.7)                 | 31(40.3)                 | 0.325           |                    |
| >64                          | 80             | 54(67.5)                 | 26(32.5)                 | 0.242           | 1.404(0.725-2.720) |
| <b>Gender</b>                |                |                          |                          |                 |                    |
| Male                         | 86             | 54(62.8)                 | 32(37.2)                 | 0.868           |                    |
| Female                       | 71             | 46(64.8)                 | 25(35.2)                 | 0.941           | 0.975(0.498-1.908) |
| <b>Family History</b>        |                |                          |                          |                 |                    |
| No                           | 143            | 92(64.3)                 | 51(35.7)                 | 0.576           |                    |
| Yes                          | 14             | 8(57.1)                  | 6(42.9)                  | 0.560           | 0.713(0.229-2.219) |
| <b>Differentiated Degree</b> |                |                          |                          |                 |                    |
| Low                          | 85             | 55(64.7)                 | 30(35.3)                 | 0.590           |                    |
| Medium                       | 60             | 39(65.0)                 | 21(35.0)                 | 0.925           | 0.967(0.476-1.964) |
| High                         | 12             | 6(50.0)                  | 6(50.0)                  | 0.126           | 0.355(0.094-1.337) |
| <b>Primary Organ</b>         |                |                          |                          |                 |                    |
| Colon                        | 75             | 45(60.0)                 | 30(40.0)                 | 0.408           |                    |
| Rectum                       | 82             | 55(67.1)                 | 27(32.9)                 | 0.228           | 1.512(0.772-2.962) |
| <b>Infiltration</b>          |                |                          |                          |                 |                    |
| Negative                     | 128            | 77(60.2)                 | 51(39.8)                 | 0.057           |                    |
| Positive                     | 29             | 23(79.3)                 | 6(20.7)                  | 0.059           | 2.545(0.966-6.701) |
| <b>Size</b>                  |                |                          |                          |                 |                    |
| <=6                          | 86             | 60(69.8)                 | 26(30.2)                 | 0.096           |                    |
| >6                           | 71             | 40(56.3)                 | 31(43.7)                 | 0.080           | 0.551(0.283-1.075) |
| <b>DFS</b>                   |                |                          |                          |                 |                    |
| 0 = progression              | 48             | 37(77.1)                 | 11(22.9)                 | <b>0.030</b>    |                    |
| 1 = death                    | 109            | 63(57.8)                 | 46(42.2)                 | <b>0.012</b>    | 0.365(0.166-0.805) |
| <b>OS</b>                    |                |                          |                          |                 |                    |
| 0 = alive                    | 58             | 44(75.9)                 | 14(24.1)                 | <b>0.017</b>    |                    |
| 1 = death                    | 99             | 56(56.6)                 | 43(43.4)                 | <b>0.008</b>    | 0.369(0.176-0.774) |

Abbreviations: 95%CI, 95% confidence interval; DFS, Disease-free survival; OS, Overall survival.

*P* values, Adjusted HR (95%CI) were assessed using Pearson  $\chi^2$  test and logistic regression analysis adjusted for age, differentiation.

**Supplementary Table 2. Correlation between DDX21 expression and clinicopathological features in the included colorectal cancer patients (*n* = 157).**

| Characteristics              | <i>n</i> = 157 | DDX21 expression          |                           | <i>P</i> -value | Adjusted OR(95%CI) |
|------------------------------|----------------|---------------------------|---------------------------|-----------------|--------------------|
|                              |                | Low (%)<br><i>n</i> = 100 | High (%)<br><i>n</i> = 57 |                 |                    |
| <b>Age</b>                   |                |                           |                           |                 |                    |
| <=64                         | 77             | 33(42.9)                  | 44(57.1)                  | 0.631           |                    |
| >64                          | 80             | 38(47.5)                  | 42(52.5)                  | 0.564           | 1.205(0.639-2.274) |
| <b>Gender</b>                |                |                           |                           |                 |                    |
| Male                         | 86             | 37(43.0)                  | 49(57.0)                  | 0.629           |                    |
| Female                       | 71             | 34(47.9)                  | 37(52.1)                  | 0.729           | 1.121(0.588-2.137) |
| <b>Family History</b>        |                |                           |                           |                 |                    |
| No                           | 143            | 68(47.6)                  | 75(52.4)                  | 0.090           |                    |
| Yes                          | 14             | 3(21.4)                   | 11(78.6)                  | 0.066           | 0.287(0.076-1.088) |
| <b>Differentiated Degree</b> |                |                           |                           |                 |                    |
| Low                          | 85             | 39(45.9)                  | 46(54.1)                  | 0.899           |                    |
| Medium                       | 60             | 26(43.3)                  | 34(56.7)                  | 0.702           | 0.876(0.446-1.723) |
| High                         | 12             | 6(50.0)                   | 6(50.0)                   | 0.977           | 0.982(0.282-3.419) |
| <b>Infiltration</b>          |                |                           |                           |                 |                    |
| Negative                     | 128            | 54(42.2)                  | 74(57.8)                  | 0.148           |                    |
| Positive                     | 29             | 17(58.6)                  | 12(41.4)                  | 0.113           | 1.941(0.856-4.401) |
| <b>Size</b>                  |                |                           |                           |                 |                    |
| <=6                          | 86             | 43(50.0)                  | 43(50.0)                  | 0.201           |                    |
| >6                           | 71             | 28(39.4)                  | 43(60.6)                  | 0.185           | 0.648(0.341-1.232) |
| <b>DFS</b>                   |                |                           |                           |                 |                    |
| 0 = progression              | 48             | 31(64.6)                  | 17(35.4)                  | 0.002           |                    |
| 1 = death                    | 109            | 40(36.7)                  | 69(63.3)                  | 0.001           | 0.287(0.139-0.595) |
| <b>OS</b>                    |                |                           |                           |                 |                    |
| 0 = alive                    | 58             | 34(58.6)                  | 24(41.4)                  | 0.013           |                    |
| 1 = death                    | 99             | 37(37.4)                  | 62(62.6)                  | 0.006           | 0.385(0.195-0.762) |

Abbreviations: 95%CI, 95% confidence interval; DFS, Disease-free survival; OS, Overall survival.

*P* values, Adjusted HR (95%CI) were assessed using Pearson  $\chi^2$  test and logistic regression analysis adjusted for age, differentiation.

**Supplementary Table 3. Short hairpin RNAs (shRNAs) sequence against ZFAS1.**

| shRNA                   | Sequence (5'-3')       |
|-------------------------|------------------------|
| ZFAS1-Homo-455          | GCCATTCTGTTCTTCGCGTCT  |
| ZFAS1-Homo-525          | GCTATTGTCTGCCGGTTAGA   |
| ZFAS1-Homo-769(shRNA-1) | GATTCAGTCTGCCTTGTAACA  |
| ZFAS1-Homo-884(shRNA-2) | CAAGGTTACTGTATAACATAGC |

**Supplementary Table 4. Short hairpin RNAs (shRNAs) against DDX21.**

| <b>shRNA</b>             | <b>Sequence (5'-3')</b> |
|--------------------------|-------------------------|
| DDX21-Homo-457(shRNA-1)  | GGAGCCTTCTGAGGAAGAAAT   |
| DDX21-Homo-933           | GCAAAGACTTCAGTGACAYCA   |
| DDX21-Homo-1245          | GGGTATTAATGTTGCCAAGA    |
| DDX21-Homo-1983(shRNA-2) | GCTCCTTGATCAACTCAAATG   |

**Supplementary Table 5. Reverse transcription polymerase chain reaction (RT-qPCR) assays.**

| <b>mRNA and LncRNA</b>   |                        |
|--|------------------------|
| <b>Relative reagents</b>   | <b>1×system (10µl)</b> |
| 5×RT Buffer  | 2 µl                   |
| Enzyme Mix   | 0.5 µl                 |
| Primer Mix   | 0.5 µl                 |
| RNA+DEPC H <sub>2</sub> O  | 7 µl (400ng)           |
| <b>Program: 37°C/15min→ 98°C/5min→ Maintain at 4°C</b>                                   |                        |
| <b>qPCR</b>  |                        |
| <b>Relative reagents</b>   | <b>1×system(10µl)</b>  |
| SYBR   | 5 µl                   |
| ROX  | 0.2 µl                 |
| Forward Primer(10µM)   | 0.4 µl                 |
| Reverse Primer(10µM)   | 0.4 µl                 |
| cDNA   | 4 µl                   |
| <b>Program: 95°C/60sec→[95°C/15sec→ 60°C/15sec→72°C/45sec]×40cycles→ Maintain at 4°C</b> |                        |

**Supplementary Table 6. Primers used in qRT-PCR assays.**

| <b>Primer</b>  | <b>Sequence (5'-3')</b>  |
|----------------|--------------------------|
| ZFAS1-Forward  | GCTATTGTCCTGCCCGTTAG     |
| ZFAS1-Reverse  | TCGTCAGGAGATCGAAGGTT     |
| DDX21-Forward  | GGCAGCGTGTCTGATTCTT      |
| DDX21-Reverse  | GCCTGCCAGAACGCTGTAGAGT   |
| POLR1A-Forward | ACAAGCCAGTTCACCGTCTTCATG |
| POLR1A-Reverse | CTGTCCTCTCCACTCTGTCCTTCC |
| POLR1B-Forward | GTGGAGTCCTCAACTACGCTGTG  |
| POLR1B-Reverse | CTGCCTCCTCATGGTGCTCAATG  |
| POLR1C-Forward | ACTATGGCTGTGGAGAACGGCTTG |
| POLR1C-Reverse | GCCTTCTTCATCTCCTTGGTTCCG |
| POLR1D-Forward | TGCCTGGCTGTCCCTCAGATCATC |
| POLR1D-Reverse | CGGTGGCGAGAGGTCTATTCTAG  |
| POLR1E-Forward | TTGTGGCTGCTGCTGTCTTAACTC |
| POLR1E-Reverse | CACCTCGCACTCGGCAACAC     |
| GAPDH-Forward  | CTCTGCTCCTCCTGTTCGAC     |
| GAPDH-Reverse  | ACCAAATCCGTTGACTCCGA     |

**Supplementary Table 7. Probes used in situ hybridization (ISH) assay.**

| <b>Digoxin-labeled probe</b> | <b>Sequence (5'-3')</b>                 |
|------------------------------|---|
| ZFAS1-(1)                    | GGAACCCGTCGAGCGGTTGGTGCCTGTGAAGCGACAT   |
| ZFAS1-(2)                    | GGTTATATAAGGGAGGTTCAGGAAGCCATTCTGTTCTTG |
| ZFAS1-(3)                    | CTACAACCTTCGATCTCCTGACGAGTTATTGTTGGCCAA |

**Supplementary Table 8. Data of lncRNAs cluster in Heatmap analysis.**

| <b>Up-regulated</b> |                    | <b>Down-regulated</b> |                    |
|---------------------|--------------------|-----------------------|--------------------|
| <b>LncRNA</b>       | <b>Fold Change</b> | <b>LncRNA</b>         | <b>Fold Change</b> |
| <b>ZFAS1</b>        | <b>6.65</b>        | RNF138P1              | 2.21               |
| TERC                | 3.17               | PSMD6-AS2             | 2.64               |
| SNORD12C            | 5.71               | PGM5P2                | 2.64               |
| SNORD12B            | 7.42               | NPY6R                 | 4.92               |
| SNORD123            | 2.39               | NCF1C                 | 2.02               |
| SNORD12             | 9.02               | NCF1B                 | 2.27               |
| SNORA71D            | 5.83               | LOC285972             | 2.37               |
| SNORA71C            | 12.02              | LOC285878             | 4.13               |
| SNORA71B            | 3.27               | LINC00675             | 2.64               |
| SNORA71A            | 17.16              | LINC00641             | 3.06               |
| SNORA65             | 4.82               | LINC00294             | 2.24               |
| SNORA31             | 3.90               | IGKV2-29              | 8.77               |
| SNORA22             | 4.87               | GVINP1                | 3.54               |
| SNORA21             | 3.98               | FENDRR                | 4.01               |
| SNORA15             | 2.10               | FENDRR                | 4.01               |
| SNORA10             | 2.24               | FAM83H-AS1            | 2.16               |
| SNHG8               | 3.45               | DLEU1                 | 2.34               |
| SNHG17              | 2.11               | BACE2-IT1             | 3.05               |
| SCARNA9L            | 3.11               | ANKRD36BP2            | 3.31               |
| SCARNA23            | 3.12               | ADAMTS9AS2            | 3.18               |
| RPL13AP20           | 2.06               | ABCC13                | 2.12               |
| RNU5F-1             | 2.13               |                       |                    |
| RNU105A             | 2.25               |                       |                    |
| PROX1-AS1           | 2.29               |                       |                    |
| MGC32805            | 2.78               |                       |                    |
| LOC541471           | 2.56               |                       |                    |
| LINC00493           | 2.10               |                       |                    |
| LINC00273           | 2.46               |                       |                    |
| HSP90AB3P           | 2.18               |                       |                    |
| HSD17B7P2           | 2.14               |                       |                    |
| FEZF1-AS1           | 3.69               |                       |                    |
| CRNDE               | 3.88               |                       |                    |
| CMAHP               | 6.25               |                       |                    |

**Supplementary Table 9. Data of mRNAs cluster in Heat map analysis.**

| Up-regulated mRNA |             | Down-regulated mRNA |             |
|-------------------|-------------|---------------------|-------------|
| mRNA              | Fold Change | mRNA                | Fold Change |
| ABCE1             | 2.81        | ABCG2               | 14.70       |
| AGO2              | 2.02        | AGL                 | 2.71        |
| BTF3              | 1.86        | AGPAT9              | 3.57        |
| CMSS1             | 2.73        | BTC                 | 2.27        |
| DCTD              | 1.54        | BTNL8               | 5.01        |
| DDIT4             | 2.21        | CLDN8               | 4.93        |
| DDX10             | 2.46        | CLMN                | 2.97        |
| <b>DDX21</b>      | <b>6.85</b> | CNNM4               | 3.68        |
| DDX31             | 2.08        | EIF4E3              | 2.16        |
| EIF3M             | 2.52        | GUCA2A              | 4.05        |
| EIF4A3            | 1.96        | GUCA2B              | 10.52       |
| GTF3C3            | 1.54        | HSPB8               | 5.93        |
| GTPBP4            | 2.59        | ITLN1               | 3.58        |
| HSP90AB1          | 2.55        | KCNIP4              | 2.95        |
| HSPA8             | 3.43        | LARGE               | 1.50        |
| HSPD1             | 3.88        | MYL12B              | 1.71        |
| ITGA2             | 8.29        | MYLK                | 4.01        |
| KCNH8             | 4.16        | PRKAG2              | 1.77        |
| KPNA7             | 1.71        | PRKCB               | 3.08        |
| LAMC2             | 6.64        | ZNF625              | 1.72        |
| LARP1             | 2.14        |                     |             |
| MYC               | 7.64        |                     |             |
| PRPF40A           | 1.65        |                     |             |
| PRPF6             | 1.89        |                     |             |
| RPL29             | 2.38        |                     |             |
| RPL30             | 2.18        |                     |             |
| RPL31             | 1.78        |                     |             |
| XRN2              | 3.63        |                     |             |
| YEATS2            | 1.64        |                     |             |
| ZC3H8             | 2.08        |                     |             |

**Supplementary Table 10. Multivariate Cox regression analysis these indicators for DFS and OS in this included colorectal cancer patients (*n* = 157).**

| Variables           | DFS               |                        |                       |              | OS                |                        |                       |              |
|---------------------|-------------------|------------------------|-----------------------|--------------|-------------------|------------------------|-----------------------|--------------|
|                     | Total<br><i>n</i> | Events<br><i>n</i> (%) | Adjusted<br>HR(95%CI) | <i>P</i>     | Total<br><i>n</i> | Events<br><i>n</i> (%) | Adjusted<br>HR(95%CI) | <i>P</i>     |
| <b>All patients</b> |                   |                        |                       |              |                   |                        |                       |              |
| <b>ZFAS1</b>        |                   |                        |                       |              |                   |                        |                       |              |
| Low                 | 100               | 63(63.0)               | 1(reference)          | -            | 100               | 56(56.0)               | 1(reference)          | -            |
| High                | 57                | 46(80.7)               | 1.936(1.301-2.880)    | <b>0.001</b> | 57                | 43(75.4)               | 1.976(1.306-2.990)    | <b>0.001</b> |
| <b>DDX21</b>        |                   |                        |                       |              |                   |                        |                       |              |
| Low                 | 71                | 40(56.3)               | 1(reference)          | -            | 71                | 37(52.1)               | 1(reference)          | -            |
| High                | 86                | 69(80.2)               | 1.671(1.128-2.475)    | <b>0.010</b> | 86                | 62(72.1)               | 1.587(1.053-2.390)    | <b>0.027</b> |
| <b>POLR1B</b>       |                   |                        |                       |              |                   |                        |                       |              |
| Low                 | 75                | 43(57.3)               | 1(reference)          | -            | 75                | 38(50.7)               | 1(reference)          | -            |
| High                | 82                | 66(80.5)               | 1.931(1.309-2.847)    | <b>0.001</b> | 82                | 61(74.4)               | 1.923(1.278-2.893)    | <b>0.002</b> |

Abbreviations: 95%CI, 95% confidence interval; DFS, Disease-free survival; OS, Overall survival.

*P* values, Adjusted HR (95%CI) were assessed using multivariate Cox regression analysis adjusted for age, differentiation.

**Supplementary Table 11. Data of mRNAs cluster of target genes in Heat map analysis.**

| Up-regulated mRNA | Fold Change | Down-regulated mRNA | Fold Change |
|-------------------|-------------|---------------------|-------------|
| ACOT9             | 1.76        |                     |             |
| ACTG1             | 1.84        |                     |             |
| AP3M2             | 2.09        |                     |             |
| APIP              | 2.11        |                     |             |
| CCT7              | 2.47        |                     |             |
| ESM1              | 4.08        | ACACB               | 3.42        |
| GLRX3             | 2.05        | APC                 | 1.54        |
| LYAR              | 2.48        | CD3G                | 2.36        |
| LYPLA1            | 2.51        | CD96                | 3.68        |
| MACC1             | 7.44        | EXT1                | 1.66        |
| MAK16             | 2.90        | FABP2               | 5.26        |
| NUP155            | 4.23        | FAM162A             | 2.52        |
| NUP35             | 1.78        | GLTP                | 3.08        |
| OLA1              | 2.25        | GNA13               | 2.02        |
| <b>POLR1B</b>     | <b>2.73</b> | GNAO1               | 10.23       |
| POLR1C            | 1.54        | MALL                | 10.24       |
| POLR2G            | 1.56        | NUMB                | 1.79        |
| POLR2K            | 2.21        | NXPE1               | 5.56        |
| POLR3F            | 1.67        | SORCS1              | 1.68        |
| RPL19             | 1.63        | SOS2                | 1.97        |
| RPL28             | 1.71        | SPIB                | 1.90        |
| RPL29             | 2.38        | TTLL7               | 1.89        |
| RPL30             | 2.18        | UGP2                | 4.15        |
| SOX9              | 7.29        | ZNF404              | 1.58        |
| TTI1              | 2.24        | ZZEF1               | 3.50        |
| TUBG1             | 1.83        |                     |             |
| WDR75             | 2.80        |                     |             |
| YEATS2            | 1.64        |                     |             |
| ZC3H8             | 2.08        |                     |             |
| ZNF280C           | 2.46        |                     |             |

**Supplementary Table 12. Correlation between POLR1B expression and clinicopathological features in the included colorectal cancer patients (*n* = 157).**

| Characteristics              | <i>n</i> = 157 | POLR1B expression         |                           | <i>P</i> -value | Adjusted OR(95%CI) |
|------------------------------|----------------|---------------------------|---------------------------|-----------------|--------------------|
|                              |                | Low (%)<br><i>n</i> = 100 | High (%)<br><i>n</i> = 57 |                 |                    |
| <b>Age</b>                   |                |                           |                           |                 |                    |
| <=64                         | 77             | 33(42.9)                  | 44(57.1)                  | 0.264           |                    |
| >64                          | 80             | 42(52.5)                  | 38(47.5)                  | 0.228           | 1.473(0.784-2.765) |
| <b>Gender</b>                |                |                           |                           |                 |                    |
| Male                         | 86             | 40(46.5)                  | 46(53.5)                  | 0.750           |                    |
| Female                       | 71             | 35(49.3)                  | 36(50.7)                  | 0.801           | 1.086(0.572-2.061) |
| <b>Family History</b>        |                |                           |                           |                 |                    |
| No                           | 143            | 69(48.3)                  | 74(51.7)                  | 0.784           |                    |
| Yes                          | 14             | 6(42.9)                   | 8(57.1)                   | 0.696           | 0.801(0.263-2.441) |
| <b>Differentiated Degree</b> |                |                           |                           |                 |                    |
| Low                          | 85             | 42(49.4)                  | 43(50.6)                  | 0.576           |                    |
| Medium                       | 60             | 26(43.3)                  | 34(56.7)                  | 0.390           | 0.744(0.379-1.459) |
| High                         | 12             | 7(58.3)                   | 5(41.7)                   | 0.725           | 1.253(0.356-4.405) |
| <b>Primary Organ</b>         |                |                           |                           |                 |                    |
| Colon                        | 75             | 32(42.7)                  | 43(57.3)                  | 0.264           |                    |
| Rectum                       | 82             | 43(52.4)                  | 39(47.6)                  | 0.155           | 1.595(0.838-3.037) |
| <b>Infiltration</b>          |                |                           |                           |                 |                    |
| Negative                     | 128            | 60(46.9)                  | 68(53.1)                  | 0.684           |                    |
| Positive                     | 29             | 24(33.8)                  | 14(48.3)                  | 0.642           | 1.212(0.539-2.726) |
| <b>Size</b>                  |                |                           |                           |                 |                    |
| <=6                          | 86             | 51(59.3)                  | 35(40.7)                  | 0.002           |                    |
| >6                           | 71             | 40(56.3)                  | 31(66.2)                  | 0.002           | 0.347(0.180-0.670) |
| <b>DFS</b>                   |                |                           |                           |                 |                    |
| 0 = progression              | 48             | 32(66.7)                  | 16(33.3)                  | <b>0.002</b>    |                    |
| 1 = death                    | 109            | 43(39.4)                  | 66(60.6)                  | <b>0.001</b>    | 0.303(0.147-0.627) |
| <b>OS</b>                    |                |                           |                           |                 |                    |
| 0 = alive                    | 58             | 37(63.8)                  | 21(36.2)                  | <b>0.003</b>    |                    |
| 1 = death                    | 99             | 38(38.4)                  | 61(61.6)                  | <b>0.001</b>    | 0.325(0.163-0.647) |

Abbreviations: 95%CI, 95% confidence interval; DFS, Disease-free survival; OS, Overall survival.

*P* values, Adjusted HR (95%CI) were assessed using Pearson  $\chi^2$  test and logistic regression analysis adjusted for age, differentiation.

**Supplementary Table 13. Evaluation the interaction propensity of lncRNA ZFAS1with DDX21.**

| Protein region | RNA region | Interaction Propensity |
|----------------|------------|------------------------|
| 126-177        | 383-434    | 94.14                  |
| 501-552        | 383-434    | 91.71                  |
| 126-177        | 33-84      | 77.15                  |
| 126-177        | 108-159    | 75.33                  |
| 501-552        | 33-84      | 74.84                  |

**Supplementary Table 14. RNA probes used for RNA pull-down assays.**

| Probe           | Sequence(5'-3')               |
|-----------------|-------------------------------|
| ZFAS1-WT        | GATTTGGAAAGAGGGAGTCACCACTGGAC |
| ZFAS1-Mut       | GATTTGGTTCTGGGAGTCACCACTGGAC  |
| ZFAS1-antisense | CTAAAACCTTCTCCCTCAGTGGTGACCTG |