Supplementary

Table S1 Patient characteristics of LUAD

| Variables | Total (n=320) | Low (n=142) | High (n=178) | Р |
|----------------|---------------|-------------|--------------|-------|
| Age | | | | 0.30 |
| <66 years | 153 [48] | 73 [51] | 80 [45] | |
| ≥66 years | 167 [52] | 69 [49] | 98 [55] | |
| Gender | | | | 0.02 |
| Female | 172 [54] | 65 [46] | 107 [60] | |
| Male | 148 [46] | 77 [54] | 71 [40] | |
| Residual tumor | | | | 0.70 |
| R0 | 306 [96] | 137 [96] | 169 [95] | |
| R1/R2 | 14 [4] | 5 [4] | 9 [5] | |
| Smoking status | | | | 0.007 |
| Nonsmoker | 45 [14] | 11 [8] | 34 [19] | |
| Current | 83 [26] | 44 [31] | 39 [22] | |
| Former | 192 [60] | 87 [61] | 105 [59] | |
| T stage | | | | 0.22 |
| T1 | 108 [34] | 41 [29] | 67 [38] | |
| T2 | 173 [54] | 84 [59] | 89 [50] | |
| T3/T4 | 39 [12] | 17 [12] | 22 [12] | |
| N stage | | | | 0.09 |
| NO | 211 [66] | 86 [61] | 125 [70] | |
| N1/N2/N3 | 109 [34] | 56 [39] | 53 [30] | |
| M stage | | | | 0.41 |
| M0 | 228 [71] | 105 [74] | 123 [69] | |
| M1/MX | 92 [29] | 37 [26] | 55 [31] | |
| Radiotherapy | | | | 0.047 |
| No | 288 [90] | 122 [86] | 166 [93] | |
| Yes | 32 [10] | 20 [14] | 12 [7] | |
| Chemotherapy | | | | 0.99 |
| No | 213 [67] | 94 [66] | 119 [67] | |
| Yes | 107 [33] | 48 [34] | 59 [33] | |

Data are presented as n [%]. LUAD, lung adenocarcinoma.

 ${\bf Table \ S2} \ {\rm Regression \ coefficient \ of \ feature \ in \ LR \ model}$

| Features | Estimate | Std. error | z value | Pr(> z) |
|--|--------------|-------------|--------------|-------------|
| (Intercept) | -0.160642518 | 0.334625359 | -0.480066777 | 0.631179911 |
| Original_shape_Flatness | -1.076946783 | 0.410015712 | -2.626598815 | 0.008624295 |
| Original_glszm_SizeZoneNonUniformity | -1.915813063 | 0.74581873 | -2.568738201 | 0.010206954 |
| Original_firstorder_Minimum | -0.894156152 | 0.781893626 | -1.143577748 | 0.252798794 |
| Original_gldm_LargeDependenceHighGrayLevelEmphasis | 1.328802785 | 0.660241068 | 2.012602443 | 0.04415647 |
| Original_glcm_ClusterProminence | 0.266224178 | 0.587584581 | 0.453082308 | 0.650489474 |
| Original_glszm_LowGrayLevelZoneEmphasis | -0.218082501 | 0.503235684 | -0.433360566 | 0.664752844 |
| Original_ngtdm_Strength | 0.150126087 | 0.371671128 | 0.403921844 | 0.686270193 |
| Original_glrlm_RunVariance | 0.43439711 | 0.831935245 | 0.522152551 | 0.60156412 |

LR, logistic regression; Std. error, standard error.

| Variables | Total (n=63) | Low (n=27) | High (n=36) | Р |
|-------------------------|--------------|------------|-------------|------|
| Age | | | | 0.21 |
| <66 years | 18 [29] | 5 [19] | 13 [36] | |
| ≥66 years | 45 [71] | 22 [81] | 23 [64] | |
| Gender | | | | 0.84 |
| Female | 16 [25] | 6 [22] | 10 [28] | |
| Male | 47 [75] | 21 [78] | 26 [72] | |
| Histology | | | | 0.76 |
| Adenocarcinoma | 49 [78] | 20 [74] | 29 [81] | |
| Squamous cell carcinoma | 14 [22] | 7 [26] | 7 [19] | |
| Smoking status | | | | 0.23 |
| Nonsmoker | 9 [14] | 6 [22] | 3 [8] | |
| Current | 18 [29] | 8 [30] | 10 [28] | |
| Former | 36 [57] | 13 [48] | 23 [64] | |
| T stage | | | | 1.00 |
| Tis/T1 | 31 [49] | 13 [48] | 18 [50] | |
| T2/T3/T4 | 32 [51] | 14 [52] | 18 [50] | |
| N stage | | | | 0.03 |
| N0 | 51 [81] | 18 [67] | 33 [92] | |
| N1/N2 | 12 [19] | 9 [33] | 3 [8] | |
| M stage | | | | 1.00 |
| M0 | 60 [95] | 26 [96] | 34 [94] | |
| M1 | 3 [5] | 1 [4] | 2 [6] | |
| KRAS mutation status | | | | 0.40 |
| Mutant | 12 [19] | 3 [11] | 9 [25] | |
| Unknown | 10 [16] | 5 [19] | 5 [14] | |
| Wildtype | 41 [65] | 19 [70] | 22 [61] | |
| EGFR mutation status | | | | 0.66 |
| Mutant | 13 [21] | 4 [15] | 9 [25] | |
| Unknown | 10 [16] | 5 [19] | 5 [14] | |
| Wildtype | 40 [63] | 18 [67] | 22 [61] | |
| Radiotherapy | | | | 0.08 |
| No | 57 [90] | 22 [81] | 35 [97] | |
| Yes | 6 [10] | 5 [19] | 1 [3] | |
| Chemotherapy | | | | 0.20 |
| | | | | |

| Data are presented as n [%] or mean ± standard deviation. OS, overall survival; TCIA-GEO, The Cancer Imaging Archive-Gene Expression | | | | |
|--|--------------|-------------|----------|------|
| OS time, mean ± SD | 39.63 ±22.03 | 33.79±22.73 | 44±20.73 | 0.07 |
| 1 | 24 [38] | 14 [52] | 10 [28] | |
| 0 | 39 [62] | 13 [48] | 26 [72] | |
| OS | | | | 0.09 |
| Yes | 17 [27] | 10 [37] | 7 [19] | |
| No | 46 [73] | 17 [63] | 29 [81] | |

Omnibus.

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Figure S1 Clinical Characteristics and the Level of Gene Expression in LUAD. (A) ALOX5 expression in normal and tumor tissues. (B) Kaplan-Meier curve for ALOX5 expression and Overall Survival (OS) in LUAD. (C) Univariate Cox and Multivariate Cox regression analyses in LUAD. ***, P<0.001. ALOX5, arachidonic acid 5-lipoxygenase; CI, confidence interval; FPKM, fragments per kilobase of transcript per million mapped reads; HR, hazard ratio; LUAD, lung adenocarcinoma; OS, overall survival.



Figure S2 Subgroup analysis and differential gene analysis of iron death-related gene set. (A) Subgroup analysis in NSCLC. (B) Differential analysis of the iron death-related gene set between high and low ALOX5 groups in LUAD. *, P<0.05; **, P<0.01; ***, P<0.001. NS, no significance; ALOX5, arachidonic acid 5-lipoxygenase; CI, confidence interval; FPKM, fragments per kilobase of transcript per million mapped reads; HR, hazard ratio; NSCLC, non-small cell lung cancer.



Figure S3 SVM model development and assessment. (A) The importance of the selected features in the SVM model. (B) ROC curve analysis of the SVM model. (C) Cross-validation ROC curve of the SVM model. (D) SVM model calibration curve evaluation. (E) DCA of the SVM model. (F) Analysis of differences between model groups. ****, P<0.0001. ALOX5, arachidonic acid 5-lipoxygenase; AUC, area under the curve; DCA, decision curve analysis; FPR, false positive rate; HL, Hosmer-Lemeshow; ROC, receiver operating characteristic; SVM, Support Vector Machine; TPR, true positive rate.