

Tables S1. Validation analyses of the ROC analyses

| A. Logistic regression analysis | | | | | | | | |
|--|----------------------|--------------------|--------------------|-----------------|--------------|---------------------|----------------|------------------|
| | | | Univariate | | | Multivariate | | |
| Variable | Cut off level | Odds ratio | 95%CI | P value | RR | 95%CI | p value | |
| Biomarkers at admission | | | | | | | | |
| CRP (mg/dL) | 8.0 | 44.6 | 9.1–336.4 | <0.001 * | 33.2 | 3.4–874.8 | 0.007 * | |
| Interleukin-6 (pg/mL) | 133.0 | 67.5 | 11.2–596.4 | <0.001 * | 38.7 | 2.6–216.1 | 0.014 | |
| KL-6 (IU/mL) | 382.0 | 28.5 | 5.1–184.4 | <0.001 * | 64.6 | 3.8–2389.5 | 0.006 * | |
| Biomarkers at 3 days after admission | | | | | | | | |
| Platelet counts (×10 ⁴ /uL) | 17.2 | 13.9 | 2.1–275.9 | 0.020* | 18.6 | 1.7–572.9 | 0.033 * | |
| CRP (mg/dL) | 2.2 | 3.4 | 0.63–26.1 | 0.17 | 0.40 | 0.010–6.4 | 0.52 | |
| HMGB-1 (ng/mL) | 13.4 | 34184160.6 | 0–NA | 0.99 | 1303873456.3 | 0–NA | 0.99 | |
| D-dimmer (ug/mL) | 1.9 | 7.5 | 1.2–47.2 | 0.025 * | 2.0 | 0.13–32.0 | 0.61 | |
| B. ROC analysis with 5 repeated 10-fold cross-validation. | | | | | | | | |
| | AUC | Sensitivity | Specificity | Accuracy | Kappa | Precision | Recall | F measure |
| Biomarkers at admission | 0.92 | 0.99 | 0.67 | 0.97 | 0.65 | 0.98 | 0.99 | 0.99 |
| Biomarkers at 3 days | 0.88 | 0.96 | 0.47 | 0.92 | 0.34 | 0.96 | 0.96 | 0.95 |
| after admission | | | | | | | | |

KL-6, Krebs von den Lungen; HMGB-1, high mobility group box 1; ROC, Receiver Operating Characteristic; 95%CI, 95% confidence interval. * A value of $p < 0.05$ was considered to represent statistical significance.

Tables S2. Correlation among biomarkers

| A. Analysis of biomarkers at admission | | | | | | | | | | | | |
|---|-----------|-----------|----------|-----------|-----------|-----------|------------|-----------|-----------|-----------|----------|-----------|
| | | | | | | | R, p value | | | | | |
| | Neut | Lym | Plt | CRP | LDH | Fer | IL-6 | IL-18 | KL-6 | HMGB-1 | sCD163 | D-dimmer |
| Neutrophil (uL) | - | 0.034 | 0.19 | 0.49 | 0.38 | 0.30 | 0.24 | 0.073 | 0.25 | 0.55 | 0.052 | 0.32 |
| | | 0.69 | 0.024 * | <0.0001 * | <0.0001 * | 0.0006 * | 0.0077 * | 0.44 | 0.0070 * | <0.0001 * | 0.59 | 0.0001 * |
| Lymphocyte (uL) | 0.034 | - | 0.019 | -0.24 | -0.19 | -0.12 | -0.26 | -0.13 | 0.054 | 0.15 | 0.047 | -0.17 |
| | 0.69 | | 0.025 * | 0.0053 * | 0.026 * | 0.18 | 0.0037 * | 0.16 | 0.56 | 0.11 | 0.62 | 0.046 * |
| Platelet (×10 ⁴ /uL) | 0.19 | 0.019 | - | -0.13 | -0.060 | -0.20 | -0.30 | -0.17 | -0.087 | 0.23 | -0.10 | 0.031 |
| | 0.024 * | 0.025 * | | 0.13 | 0.49 | 0.020 * | 0.0006 * | 0.076 | 0.35 | 0.014 * | 0.30 | 0.72 |
| CRP (mg/dL) | 0.49 | -0.24 | -0.13 | - | 0.68 | 0.71 | 0.64 | 0.47 | 0.27 | 0.17 | 0.24 | 0.37 |
| | <0.0001 * | 0.0053 * | 0.13 | | <0.0001 * | <0.0001 * | <0.0001 * | <0.0001 * | 0.0033 * | 0.084 | 0.012 * | <0.0001 |
| Lactate dehydrogenase (IU/L) | 0.38 | -0.19 | -0.060 | 0.68 | - | 0.69 | 0.53 | 0.57 | 0.37 | 0.31 | 0.29 | 0.40 |
| | <0.0001 * | 0.026 * | 0.49 | <0.0001 * | | <0.0001 * | <0.0001 * | <0.0001 * | <0.0001 * | 0.0011 * | 0.0021 * | <0.0001 * |
| Ferritin (ng/mL) | 0.30 | -0.12 | -0.20 | 0.71 | 0.69 | - | 0.63 | 0.60 | 0.31 | 0.15 | 0.31 | 0.21 |
| | 0.0006 * | 0.18 | 0.020 * | <0.0001 * | <0.0001 * | | <0.0001 * | <0.0001 * | 0.0005 * | 0.13 | 0.0009 * | 0.020 * |
| Interleukin-6 (pg/mL) | 0.24 | -0.26 | -0.30 | 0.64 | 0.53 | 0.63 | - | 0.48 | 0.21 | -0.0049 | 0.28 | 0.16 |
| | 0.0077 * | 0.0037 * | 0.0006 * | <0.0001 * | <0.0001 * | <0.0001 * | | <0.0001 * | 0.021 * | 0.96 | 0.0028 * | 0.11 |
| Interleukin-18 (pg/mL) | 0.073 | -0.13 | -0.17 | 0.47 | 0.57 | 0.60 | 0.48 | - | 0.13 | 0.16 | 0.35 | 0.16 |
| | 0.44 | 0.16 | 0.076 | <0.0001 * | <0.0001 * | <0.0001 * | <0.0001 * | | 1.15 | 0.11 | 0.0003 * | 0.11 |
| KL-6 (IU/mL) | 0.25 | 0.054 | -0.087 | 0.27 | 0.37 | 0.31 | 0.21 | 0.13 | - | 0.19 | 0.28 | 0.35 |
| | 0.0070 * | 0.56 | 0.35 | 0.0033 * | <0.0001 * | 0.0005 * | 0.021 * | 1.15 | | 0.060 * | 0.038 * | <0.0001 * |
| HMGB-1 (ng/mL) | 0.55 | 0.15 | 0.23 | 0.17 | 0.31 | 0.15 | -0.0049 | 0.16 | 0.19 | - | 0.086 | 0.11 |
| | <0.0001 * | 0.11 | 0.014 * | 0.084 | 0.0011 * | 0.13 | 0.96 | 0.11 | 0.060 * | | 0.37 | 0.25 |
| Soluble CD163 (ng/mL) | 0.052 | 0.047 | -0.10 | 0.24 | 0.29 | 0.31 | 0.28 | 0.35 | 0.28 | 0.086 | - | |
| | 0.59 | 0.62 | 0.30 | 0.012 * | 0.0021 * | 0.0009 * | 0.0028 * | 0.0003 * | 0.038 * | 0.37 | | |
| D-dimer (ug/mL) | 0.32 | -0.17 | 0.031 | 0.37 | 0.40 | 0.21 | 0.16 | 0.16 | 0.35 | 0.11 | 0.32 | - |
| | 0.0001 * | 0.046 * | 0.72 | <0.0001 | <0.0001 * | 0.020 * | 0.11 | 0.11 | <0.0001 * | 0.25 | 0.0007 * | |
| B. Analysis of biomarkers at 3 days after admission | | | | | | | | | | | | |
| | | | | | | | R, p value | | | | | |
| | Neut | Lym | Plt | CRP | LDH | Fer | IL-6 | HMGB-1 | sCD163 | D-dimmer | | |
| Neutrophil (uL) | - | -0.19 | 0.28 | 0.44 | 0.42 | 0.35 | 0.11 | 0.69 | 0.16 | 0.34 | | |
| | | 0.039 * | 0.0021 * | <0.0001 * | <0.0001 * | 0.0006 * | 0.29 | <0.0001 * | 0.12 | 0.0007 * | | |
| Lymphocyte (uL) | -0.19 | - | 0.29 | -0.044 | -0.48 | -0.45 | -0.47 | -0.21 | -0.17 | -0.27 | | |
| | 0.039 * | | 0.0011 * | 0.63 | <0.0001 * | <0.0001 * | <0.0001 * | 0.049 * | 0.11 | 0.0080 * | | |
| Platelet (×10 ⁴ /uL) | 0.28 | 0.29 | - | -0.044 | -0.070 | -0.092 | -0.31 | 0.079 | -0.0088 | 0.11 | | |
| | 0.0021 * | 0.0011 * | | 0.63 | 0.44 | 0.38 | 0.0024 * | 0.46 | 0.93 | 0.28 | | |
| CRP (mg/dL) | 0.44 | -0.044 | -0.044 | - | 0.69 | 0.67 | 0.56 | 0.43 | 0.36 | 0.37 | | |
| | <0.0001 * | 0.63 | 0.63 | | <0.0001 * | <0.0001 * | <0.0001 * | <0.0001 * | 0.0005 * | 0.0002 * | | |
| Lactate dehydrogenase (IU/L) | 0.42 | -0.48 | -0.070 | 0.69 | - | 0.77 | 0.51 | 0.54 | 0.42 | 0.36 | | |
| | <0.0001 * | <0.0001 * | 0.44 | <0.0001 * | | <0.0001 * | <0.0001 * | <0.0001 * | <0.0001 * | 0.0004 * | | |
| Ferritin (ng/mL) | 0.35 | -0.45 | -0.092 | 0.67 | 0.77 | - | 0.54 | 0.50 | 0.45 | 0.29 | | |
| | 0.0006 * | <0.0001 * | 0.38 | <0.0001 * | <0.0001 * | | <0.0001 * | <0.0001 * | <0.0001 * | 0.013 * | | |

| | | | | | | | | | | | | |
|-----------------------|-----------|-----------|----------|-----------|-----------|-----------|----------|---------|-----------|-----------|--|--|
| Interleukin-6 (pg/mL) | 0.11 | -0.47 | -0.31 | 0.56 | 0.51 | 0.54 | - | 0.19 | 0.27 | 0.022 | | |
| | 0.29 | <0.0001 * | 0.0024 * | <0.0001 * | <0.0001 * | <0.0001 * | | 0.073 | 0.0092 * | 0.85 | | |
| HMGB-1 (ng/mL) | 0.69 | -0.21 | 0.079 | 0.43 | 0.54 | 0.50 | 0.19 | - | 0.086 | 0.27 | | |
| | <0.0001 * | 0.049 * | 0.46 | <0.0001 * | <0.0001 * | <0.0001 * | 0.073 | | 0.37 | 0.018 * | | |
| Soluble CD163 (ng/mL) | 0.16 | -0.17 | -0.0088 | 0.36 | 0.42 | 0.45 | 0.27 | 0.086 | - | 0.47 | | |
| | 0.12 | 0.11 | 0.93 | 0.0005 * | <0.0001 * | <0.0001 * | 0.0092 * | 0.37 | | <0.0001 * | | |
| D-dimer (ug/mL) | 0.34 | -0.27 | 0.11 | 0.37 | 0.36 | 0.29 | 0.022 | 0.27 | 0.47 | - | | |
| | 0.0007 * | 0.0080 * | 0.28 | 0.0002 * | 0.0004 * | 0.013 * | 0.85 | 0.018 * | <0.0001 * | | | |

Neut, neutrophil; Lym, lymphocyte; Plt, platelet; LDH, lactate dehydrogenase; Fer, ferritin; IL-6, interleukin-6; IL-18, interleukin-18; sCD163, soluble CD163. * A value of $p < 0.05$ was considered to represent statistical significance.