

Figure S1 (A) The distribution of patients in 17 participating hospitals (n=527). (B) The distribution of low-dose corticosteroid and control groups in different hospitals (n=289). (C) The distribution of the low-dose corticosteroid group in different hospitals (n=65).



Figure S2 Intubation rate curves for the low-dose and control groups within a 14-day period in the matched sample (A) and original sample (B).

| Table S1 Distribution of acute respiratory distress syndrome risk factors in the low-dose and control groups. There was no significant different | ce |
|--|----|
| between two groups (P=0.433) | |

| Risk factor | Low-dose corticosteroid (n=65), n (%) | Non-corticosteroid (n=224), n (%) |
|-----------------------|---------------------------------------|-----------------------------------|
| Pneumonia | 52 (81.3) | 167 (74.6) |
| Aspiration | 5 (7.8) | 7 (3.1) |
| Drowning | 0 | 1 (0.4) |
| Pulmonary contusion | 1 (1.6) | 6 (2.7) |
| Trauma | 0 | 5 (2.2) |
| Extrapulmonary sepsis | 3 (4.7) | 18 (8.0) |
| Pancreatitis | 1 (1.6) | 10 (4.5) |
| Others | 2 (3.1) | 10 (4.5) |
| Missing data | 1 (1.5) | 0 |

Table S2Aetiological diagnoses in patients with pneumonia-related acute respiratory distress syndrome in the low-dose and control groups.There was no significant difference between two groups (P=0.574)

| Pathogen | Low-dose corticosteroid (n=52), n (%) | Non-corticosteroid (n=167), n (%) |
|-----------------------|---------------------------------------|-----------------------------------|
| Gram- bacillus | 3 (5.7) | 15 (8.9) |
| Gram+ coccus | 0 | 1 (0.6) |
| Fungus | 1 (1.9) | 6 (3.6) |
| Influenza virus | 10 (19.2) | 38 (22.8) |
| Other viruses | 2 (3.9) | 2 (1.2) |
| Pneumocystis | 2 (3.9) | 1 (0.6) |
| Tuberculosis bacillus | 0 | 1 (0.6) |
| Atypical pathogens | 0 | 4 (2.4) |
| Mixed infection | 5 (9.6) | 20 (12.0) |
| Unknown | 29 (55.8) | 79 (47.3) |

 Table S3 Comparison of baseline characteristics between high-dose and control groups

| Variable | High-dose corticosteroid (n=65) Non-corticosteroid (n=224) | | P value |
|--|--|--------------------|---------|
| Male sex, n (%) | 45 (69.2) 156 (69.6) | | 0.949 |
| Age, median (IQR), years | 58.0 (44.0–70.0) | 57.0 (45.0–69.0) | 0.874 |
| BMI, median (IQR) | 24.2 (22.0–26.2) | 24.2 (21.5–26.7) | 0.776 |
| PFR at admission (mmHg) | 107.0 (80.0–162.8) | 115.2 (84.3–162.0) | 0.695 |
| APACHE II score, median (IQR) | 17 (10–23) | 15 (10–21) | 0.240 |
| SOFA score, median (IQR) | 7 (4–9) | 6 (4–10) | 0.972 |
| Intrapulmonary ARDS, n (%) | 60 (92.3) | 182 (81.3) | 0.033 |
| Underlying disease condition, n (%) | | | |
| Hypertension | 24 (36.9) | 65 (29.3) | 0.241 |
| Diabetes mellitus | 14 (21.5) | 43 (19.4) | 0.700 |
| Chronic cardiac insufficiency | 3 (4.6) | 8 (3.6) | 0.716 |
| Chronic kidney disease | 6 (9.2) | 20 (9.0) | 0.956 |
| Immunosuppression* | 32 (49.2) | 39 (17.4) | 0.000 |
| Laboratory test results at ICU admission | | | |
| D0 WBC, median (IQR) (×10 ⁹ /L) | 11.3 (8.9–16.4) | 10.0 (6.3–15.3) | 0.150 |
| D0 PCT, median (IQR) (ng/mL) | 1.1(0.3–5.0) | 2.0 (0.4–10.5) | 0.077 |
| D0 CRP, median (IQR) (mg/L) | 130.1 (60.7–211.7) | 124.7 (42.4–200.0) | 0.648 |
| D0 lactic acid, median (IQR) (mmol/L) | 1.9 (1.2–2.7) | 1.8 (1.1–2.8) | 0.905 |

*, immunosuppression was defined as a haematologic malignancy or a solid tumour; or administration of steroids or any immunosuppressive drug within a month; or administration of radiation therapy or chemotherapy within a year (Same as *Table 1*).

Table S4 Comparison of baseline characteristics between other dose corticosteroid group and control groups

| Variable | Other-dose corticosteroid (n=189) | Non-corticosteroid (n=224) | P value |
|--|-----------------------------------|----------------------------|---------|
| Male sex, n (%) | 137 (72.5) | 156 (69.6) | 0.526 |
| Age, median (IQR), years | 55.0 (43.0–68.0) | 57.0 (45.0–69.0) | 0.666 |
| BMI, median (IQR) | 24.0 (21.7–26.1) | 24.2 (21.5–26.7) | 0.458 |
| PFR at admission (mmHg) | 110.5 (74.0–160.0) | 113.0 (84.1–160.1) | 0.310 |
| APACHE II score, median (IQR) | 18.0 (13.0–23.5) | 15.0 (10.0-21.0) | 0.002 |
| SOFA score, median (IQR) | 8.0 (5.0–11.0) | 6.0 (4.0–10.0) | 0.002 |
| Intrapulmonary ARDS, n (%) | 158 (83.6) | 182 (81.3) | 0.533 |
| Underlying disease condition, n (%) | | | |
| Hypertension | 70 (37.0) | 65 (29.3) | 0.095 |
| Diabetes mellitus | 39 (20.7) | 43 (19.4) | 0.729 |
| Chronic cardiac insufficiency | 12 (6.3) | 8 (3.6) | 0.197 |
| Chronic kidney disease | 22 (11.7) | 20 (9.0) | 0.370 |
| Immunosuppression* | 67 (35.4) | 39 (17.4) | 0.000 |
| Laboratory test results at ICU admission | | | |
| D0 WBC, median (IQR) (×10 ⁹ /L) | 10.6 (5.7–13.9) | 10.4 (6.3–15.3) | 0.910 |
| D0 PCT, median (IQR) (ng/mL) | 0.9 (0.3–5.0) | 2.0 (0.4–10.5) | 0.052 |
| D0 CRP, median (IQR) (mg/L) | 107.8 (39.3–180.4) | 124.7 (46.8–200.0) | 0.114 |
| D0 lactic acid, median (IQR) (mmol/L) | 1.9 (1.3–2.8) | 1.8 (1.1–2.8) | 0.468 |

*, immunosuppression was defined as a haematologic malignancy or a solid tumour; or administration of steroids or any immunosuppressive drug within a month; or administration of radiation therapy or chemotherapy within a year (Same as *Table 1*).

Table S5 Comparison of outcomes between the low-dose and control groups in the original sample

| Outcome | Low-dose corticosteroid (n=65) Non-corticosteroid (n=224) | | P value |
|--|---|------------------|---------|
| Duration of mechanical ventilation* (days) | 11.0 (7.0–14.0) | 8.0 (5.0–12.0) | 0.001 |
| Nosocomial infection, n (%) | 14 (21.5) | 59 (26.3) | 0.433 |
| New organ failure, n (%) | 27 (41.5) | 94 (42.0) | 0.951 |
| Ventilator free days at day 28, d | 14.0 (1.5–19.5) | 17.0 (1.0–27.8) | 0.188 |
| ICU length of stay (days) | 15.5 (10.0–24.0) | 10.0 (6.0–17.0) | 0.000 |
| Hospital length of stay (days) | 23.0 (16.0–36.3) | 17.0 (10.0–26.0) | 0.000 |
| ICU mortality, n (%) | 28 (43.1) | 87 (38.8) | 0.539 |
| Hospital mortality, n (%) | 29 (44.6) | 91 (40.6) | 0.565 |

*, Only patients with intubation were included.

Table S6 Comparison of outcomes between the high-dose and control groups

| Outcome | High-dose corticosteroid (n=41) | Non-corticosteroid (n=224) | P value |
|--|---------------------------------|----------------------------|---------|
| Duration of mechanical ventilation* (days) | 10.0 (4.0–14.0) | 10.0 (5.0–13.6) | 0.378 |
| Nosocomial infection, n (%) | 9 (22.0) | 59 (26.3) | 0.554 |
| New organ failure, n (%) | 15 (36.6) | 94 (42.0) | 0.520 |
| Ventilator free days at day 28 (days) | 16.0 (0.5–28.0) | 17.0 (1.0–27.3) | 0.581 |
| ICU length of stay (days) | 14.0 (7.5–35.0) | 10.0 (6.0–17.0) | 0.001 |
| Hospital length of stay (days) | 23.0 (12.5–26.0) | 17.0 (10.0–26.0) | 0.002 |
| ICU mortality, n (%) | 16 (39.0) | 87 (38.8) | 0.982 |
| Hospital mortality, n (%) | 17 (41.5) | 91 (40.6) | 0.920 |

*, Only patients with intubation were included.

Table S7 Comparison of outcomes between the other dose corticosteroid group and control groups

| Outcome | Other dose corticosteroid (n=189) | Non-corticosteroid (n=224) | P value |
|--|-----------------------------------|----------------------------|---------|
| Duration of mechanical ventilation* (days) | 7.0 (3.0–11.0) | 5.0 (0.0–10.0) | 0.024 |
| Nosocomial infection, n (%) | 49 (25.9) | 59 (26.3) | 0.924 |
| New organ failure, n (%) | 101 (53.4) | 94 (42.0) | 0.020 |
| Ventilator free days at day 28 (days) | 4.0 (0.0–21.0) | 17.0 (1.0–27.8) | 0.000 |
| ICU length of stay (days) | 11.0 (6.0–21.0) | 10.0 (6.0–17.0) | 0.063 |
| Hospital length of stay (days) | 17.0 (8.8–30.3) | 17.0 (10.0–26.0) | 0.444 |
| ICU mortality, n (%) | 96 (50.8) | 87 (38.8) | 0.015 |
| Hospital mortality, n (%) | 101 (53.4) | 91 (40.6) | 0.009 |

*, Only patients with intubation were included.

| Table S8. | Univariate | Cox r | regression | analysis | for | factors | associated |
|-----------|------------|---------|-------------|-----------|-------|---------|------------|
| | with hos | pital r | mortality i | n origina | ıl sa | mple | |

| Variable | HR (95% CI) | p value | |
|-------------------------|-----------------|---------|--------------|
| Age≥65 years | 1.71(1.19-2.46) | .004 | ⊢ → |
| Male | 1.06(0.71-1.59) | .763 | ⊢●1 |
| SOFA≥6 | 1.66(1.11-2.49) | .013 | ⊢ _●i |
| ARDS Severity | 1.27(0.95-1.71) | .108 | ⊢●→ |
| Intrapulmonary ARDS | 1.30(0.77-2.21) | .326 | ⊢ ●1 |
| Immunosuppression | 1.68(1.15-2.44) | .007 | ⊢ |
| Low-dose corticosteroid | 0.82(0.54-1.25) | .348 | ⊢●─┤ |
| | | | 0 1 2 3 |

 Table S9. Multivariate Cox regression analysis for factors associated with hospital mortality in original sample

| Variable | HR (95% CI) | p value | |
|-------------------------|-----------------|---------|---------------|
| Age≥65 years | 1.58(1.09-2.28) | .016 | ⊢ •−−1 |
| Male | 1.02(0.67-1.55) | .930 | F- • 1 |
| SOFA≥6 | 1.64(1.09-2.47) | .018 | ⊢ •I |
| ARDS Severity | 1.29(0.96-1.72) | .095 | ⊧ _ ●I |
| Intrapulmonary ARDS | 1.45(0.84-2.50) | .181 | ⊢ ∎1 |
| Immunosuppression | 1.58(1.05-2.39) | .029 | ⊢ ● I |
| Low-dose corticosteroid | 0.62(0.39-0.98) | .041 | ⊢●→ |
| | | | 0 1 2 3 |

Table S10 Effects of corticosteroids on mortality using multivariate Cox regression analysis in the original sample

| Subaroup | Hospital mortality | | |
|--|--------------------|-------|--|
| Subgroup | HR (95% CI) | Р | |
| All patients (n=289) | 0.63 (0.39–0.98) | 0.041 | |
| Patients with intrapulmonary ARDS (n=242) | 0.55 (0.34–0.90) | 0.017 | |
| Patients with mechanical ventilation (n=209) | 0.55 (0.34–0.91) | 0.019 | |
| Patients with shock (n=97) | 0.64 (0.35–1.18) | 0.635 | |
| Patients with influenza (n=48) | 0.27 (0.07–1.03) | 0.056 | |
| Patients without immunosuppression (n=218) | 0.41 (0.20–0.87) | 0.020 | |