

Supplementary Table S1. Demographics, comorbidities, and biometrical characteristics of critically ill COVID-19 patients.

	COVID-19 patients (n=217)
Female gender, n (%)	66 (30.4)
Age, median (min-max)	68 (32–102)
Clinical features	
BMI 25–30, n (%)	148/186* (79.5)
BMI 30–35, n (%)	31/186* (16.7)
BMI >35, n (%)	8/186* (4.3)
Comorbidities	
Hypertension, n (%)	101/215* (47)
Dyslipidaemia, n (%)	55/215* (25.6)
COPD, n (%)	10/225* (4.7)
Neoplasia, n (%)	15/215* (7)
CD, n (%)	30/215* (14)
CKD, n (%)	9/215* (4.2)
DM, n (%)	28/215* (13)
Thyroid dysfunction, n (%)	6/215* (2.8)
ICU prognostic indicators	
SOFA score (>5), n (%)	83 (38.2)
SOFA score points, median (min-max)	5 (2–11)
APACHE II score (>14), n (%)	140 (64.5)
APACHE II score points, median (min-max)	15 (3–24)
Time duration between symptoms onset and hospital admission, median (min-max)	5 (1–15) / 186*
Time duration between hospital admission and outcome, median (min-max)	20 (4–137) / 186*
Clinical characteristics	
pO ₂ on air (%), median (min-max)	56 (40.5–85) / 186*
Pulmonary embolism, n (%)	2 (0.9)
Myositis, n (%)	3 (1.4)
Cognitive dysfunction, n (%)	18 (8.3)
Laboratory findings	
Hyperferritinæmia (>250 µg/l), n (%)	154/191* (80.6)
Ferritin levels, median (min-max)	556.2 (38–25466)
D-dimers, n (%)	55/194* (28.4)
D-dimer levels, median (min-max)	1.38 (0.1–17.6)
CRP (>0.5 mg/dl), n (%)	208/214* (97.2)
CRP levels, median (min-max)	11.4 (0.3–41.1)
Creatinine (>1.4 mg/dl), n (%)	22/216* (10.2)
Creatinine levels, median (min-max)	0.9 (0.4–5.8)
HbA1c (>6.5), n (%)	34/150* (22.7)
HbA1c levels, median (min-max)	6 (4.5–13)
Treatment	
Azithromycin before hospitalisation, n (%)	39/186* (21)
Dexamethasone after intubation, n (%)	186/186* (100)

*BMI: Body Mass Index; COPD: chronic obstructive pulmonary disease; CD: cardiovascular disease; CKD: chronic kidney disease; DM: diabetes mellitus; SOFA: Sequential Organ Failure Assessment score; APACHE II: acute physiology and chronic health evaluation II; CRP: C-reactive protein; HbA1c: haemoglobin A1c.

Supplementary Table S2. Comparison between critically ill COVID-19 patients with at least one autoantibody and without.

	Patients negative in autoantibodies (n=43)	Patients positive in at least one autoantibody (n=174)	p-value
Female gender, n (%)	13 (30.2)	53 (30.5)	0.977
Age, median (min-max)	67 (47-87)	69 (32-102)	0.927
Covid-19 IgG antibodies, n (%)	28 (65.1)	150 (86.2)	0.001
Titres of Covid-19 IgG antibodies, median (min-max)	4.3 (0.1-10.1)	7.8 (0.1-11.4)	0.014
Activity of neutralising SARS-CoV-2 RBD antibodies, n (%)	33 (76.7)	159 (91.4)	0.007
Titres of neutralising SARS-CoV-2 RBD antibodies, median (min-max)	79 (12-97)	91 (4-100)	0.018
Clinical features			
Death, n (%)	23 (53.5)	83 (47.7)	0.497
Time duration between symptoms onset and hospital admission, median (min-max)	5 (1-10) / 36*	5 (1-15) / 150*	0.859
Time duration between hospital admission and outcome, median (min-max)	23 (5-137) / 36*	19 (4-120) / 150*	0.275
BMI 25-30, n (%)	29/36* (80.6)	119/150* (79.3)	0.870
BMI 30-35, n (%)	5/36* (13.9)	26/150* (17.3)	0.618
BMI >35, n (%)	2/36* (5.6)	6/150* (4)	0.653
Comorbidities			
Hypertension, n (%)	20/41* (48.8)	81 (46.6)	0.797
Dyslipidaemia, n (%)	10/41* (24.4)	45 (25.9)	0.846
COPD, n (%)	4/41* (9.8)	6 (3.4)	0.084
Neoplasia, n (%)	5/41* (12.2)	10 (5.7)	0.145
CD, n (%)	5/41* (12.2)	25 (14.4)	0.718
CKD, n (%)	2/41* (4.9)	7 (4)	0.806
DM, n (%)	8/41* (19.5)	20 (11.5)	0.170
Thyroid dysfunction, n (%)	0/41* (0)	6 (3.4)	0.598
ICU prognostic indicators			
SOFA score (>5), n (%)	14 (32.6)	69 (39.7)	0.391
SOFA score points, median (min-max)	5 (3-11)	5 (2-11)	0.673
APACHE II score (>14), n (%)	30 (69.8)	110 (63.2)	0.422
APACHE II score points, median (min-max)	15 (6-22)	15 (3-24)	0.301
Clinical characteristics			
pO ₂ on air (%), median (min-max)	57.5 (40.5-75) / 36*	55.2 (42-85) / 150*	0.037
Pulmonary embolism, n (%)	0 (0)	2 (1.1)	1
Myositis, n (%)	0 (0)	3 (1.7)	1
Cognitive dysfunction, n (%)	4 (9.3)	14 (8)	0.761
Laboratory findings			
Hyperferritinæmia (>250 µg/l), n (%)	29/37* (78.4)	125/154* (81.2)	0.700
Ferritin levels, median (min-max)	602.2 (74.9-14959)	546.9 (38-25466)	0.887
D-dimers, n (%)	15/40* (37.5)	40/154* (26)	0.150
D-dimer levels, median (min-max)	1.7 (0.4-10)	1.3 (0.1-17.6)	0.555
CRP (>0.5 mg/dl), n (%)	42 (97.7)	166/171 (97.1)	0.832
CRP levels, median (min-max)	12.5 (0.5-32)	10.3 (0.3-41.1)	0.126
Creatinine (>1.4 mg/dl), n (%)	8 (18.6)	14/173* (8.1)	0.041
Creatinine levels, median (min-max)	1 (0.4-5.8)	0.9 (0.4-4.2)	0.473
HbA1c (>6.5), n (%)	8/26* (30.8)	26/124* (21)	0.278
HbA1c levels, median (min-max)	6.1 (4.5-10.6)	6 (4.5-13)	0.611
Treatment			
Azithromycin before hospitalisation, n (%)	7/36* (19.4)	32/150* (21.3)	0.803
Dexamethasone after intubation, n (%)	36/36* (100)	150/150* (100)	1

* Available data; BMI: Body Mass Index; COPD: chronic obstructive pulmonary disease; CD: cardiovascular disease; SOFA: Sequential Organ Failure Assessment score; APACHE II: acute physiology and chronic health evaluation II; CRP: C-reactive protein; HbA1c: haemoglobin A1c.

Supplementary Table S3. Characteristics of critically ill COVID-19 patients with two consecutive blood samples available for analysis.

	Total (n=60)
Female gender, n (%)	15 (25)
Age, median (min-max)	68 (42–90)
Outcome, n (%)	30 (50)
Clinical features	
Time duration between symptoms onset and hospital admission, median (min-max)	5 (1-15)
Time duration between hospital admission and outcome, median (min-max)	42 (16–137)
BMI 25–30, n (%)	50 (83.3)
BMI 30–35, n (%)	8 (13.3)
BMI >35, n (%)	2 (3.3)
Comorbidities	
Hypertension, n (%)	31 (51.7)
Dyslipidaemia, n (%)	17 (28.3)
COPD, n (%)	5 (8.3)
Neoplasia, n (%)	4 (6.7)
CD, n (%)	12 (20)
CKD, n (%)	3 (5)
DM, n (%)	8 (13.3)
Thyroid dysfunction, n (%)	2 (3.3)
ICU prognostic indicators	
SOFA score (>5), n (%)	7 (11.7)
SOFA score points, median (min-max)	4.5 (3-6)
APACHE II score (>14), n (%)	47 (78.3)
APACHE II score points, median (min-max)	17 (11–24)
Clinical characteristics	
pO ₂ on air (%), median (min-max)	58 (42-85) / 186*
Pulmonary embolism, n (%)	1 (1.7)
Myositis, n (%)	1 (1.7)
Cognitive dysfunction, n (%)	5 (8.3)
Laboratory findings	
Hyperferritinæmia (>250 µg/l), n (%)	48/59* (81.4)
Ferritin levels, median (min-max)	490.4 (38–14959)
D-dimers, n (%)	3/59* (5.1)
D-dimer levels, median (min-max)	1 (0.1–10)
CRP (>0.5 mg/dl), n (%)	59/59* (100)
CRP levels, median (min-max)	14.2 (0.6–39)
Creatinine (>1.4 mg/dl), n (%)	12 (20)
Creatinine levels, median (min-max)	1 (0.4–2.8)
HbA1c (>6.5), n (%)	14/57* (24.6)
HbA1c levels, median (min-max)	6 (4.5–10.2)
Treatment	
Azythromycin before hospitalisation, n (%)	8 (13.3)
Dexamethasone after intubation, n (%)	60 (100)

*Available data; BMI: Body Mass Index; COPD: chronic obstructive pulmonary disease; CD: cardiovascular disease; SOFA: Sequential Organ Failure Assessment score; APACHE II: acute physiology and chronic health evaluation II; CRP: C-reactive protein; HbA1c: haemoglobin A1c.

Supplementary Table S4. Pattern and titre of antinuclear antibodies (ANA).

	Patients (n=217)	Controls (n=117)
ANA, N (%)	105 (48.4)	25 (21.4)
Type		
Fine speckled pattern, n (%)	74 (34.1)	21 (17.9)
Speckled pattern, n (%)	22 (10.1)	0 (0)
Fine speckled cytoplasmic pattern, n (%)	3 (1.4)	2 (1.7)
Speckled cytoplasmic pattern, n (%)	4 (1.8)	0 (0)
Nucleolar pattern, n (%)	1 (0.5)	2 (1.7)
Vimentin pattern, n (%)	1 (0.5)	0 (0)
Titre		
1:160, n (%)	66 (30.4)	15 (12.8)
1:320, n (%)	33 (15.2)	5 (4.3)
1:640, n (%)	6 (2.8)	5 (4.3)

Supplementary Table S5. Pattern and titre of antineutrophil cytoplasmic antibodies (ANCA).

	Patients (n=217)	Controls (n=117)
ANCA, n (%)	30 (13.8)	14 (12.0)
P-ANCA, n (%)	13 (6.0)	12 (10.3)
1:20, n (%)	4 (1.8)	2 (1.7)
1:40, n (%)	1 (0.5)	4 (3.4)
1:80, n (%)	5 (2.3)	3 (2.6)
1:160, n (%)	1 (0.5)	0 (0)
1:320, n (%)	1 (0.5)	2 (1.7)
1:640, n (%)	1 (0.5)	1 (0.9)
C-ANCA, n (%)	17 (7.8)	2 (1.7)
1:20, n (%)	5 (2.3)	0 (0)
1:40, n (%)	9 (4.1)	0 (0)
1:80, n (%)	0 (0)	1 (0.9)
1:160, n (%)	2 (0.9)	1 (0.9)
1:320, n (%)	0 (0)	0 (0)
1:640, n (%)	1 (0.5)	0 (0)

Supplementary Table S6. Myositis-related autoantibodies.

	Patients (n=203)	Controls (n=117)
Myositis-related autoantibodies, n (%)	22 (10.8)	1 (0.9)
Mi-2a, n (%)	1 (0.5)	0 (0)
Mi-2β, n (%)	0 (0)	0 (0)
TIF1γ, n (%)	3 (1.5)	0 (0)
MDA5, n (%)	1 (0.5)	0 (0)
NXP2, n (%)	1 (0.5)	0 (0)
SAE1, n (%)	0 (0)	0 (0)
Ku, n (%)	5 (2.5)	0 (0)
PM-Scl100, n (%)	2 (1)	0 (0)
PM-Scl75, n (%)	4 (2)	0 (0)
Jo-1, n (%)	1 (0.5)	0 (0)
SRP, n (%)	0 (0)	0 (0)
PL-7, n (%)	2 (1)	1 (0.9)
PL-12, n (%)	4 (2)	0 (0)
EJ, n (%)	0 (0)	0 (0)
OJ, n (%)	3 (1.5)	0 (0)

Supplementary Table S7. Titres of autoantibodies in seropositive severe COVID-19 patients.

Autoantibodies	Titres
ANA [median dilution (min-max)]	160 (80–1280)
ANCA [median dilution (min-max)]	40 (20–640)
C-ANCA [median dilution (min-max)]	40 (20–640)
P-ANCA [median dilution (min-max)]	80 (80–640)
ENAs [median intensity (min-max), low 11–25, moderate 26–50, high >50]	28 (21–72)
Myositis related autoantibodies [median intensity (min-max), low 11–25, moderate 26–50, high >50]	32 (21–116)
Anti-CCP [median I.U. (min-max), cut-off limit: 20 I.U.]	21 (31.5–468)
Anti-dsDNA [median I.U. (min-max), cut-off limit: 16 I.U.]	24 (17.6–48.4)
Anti-TPO, [median I.U. (min-max), cut-off limit: 100 I.U.]	183 (101–4099)
Anti-β2-GPI IgG [median I.U. (min-max), cut-off limit: 100 I.U.]	125.5 (101–445)
Anti-β2-GPI IgM [median I.U. (min-max), cut-off limit: 100 I.U.]	182.5 (105–1114)
Anti-CL IgG [median I.U. (min-max), cut-off limit: 100 I.U.]	240.5 (127–744)
Anti-CL IgM [median I.U. (min-max), cut-off limit: 100 I.U.]	188 (102–664)
Anti-TPO, [median I.U. (min-max), cut-off limit: 100 I.U.]	183 (101–4099)
Anti-TG, [median I.U. (min-max), cut-off limit: 0.6 I.U.]	1.2 (0.7–2.9)