Supplemental Information

Loss of BcI-6-Expressing T Follicular Helper

Cells and Germinal Centers in COVID-19

Naoki Kaneko, Hsiao-Hsuan Kuo, Julie Boucau, Jocelyn R. Farmer, Hugues Allard-Chamard, Vinay S. Mahajan, Alicja Piechocka-Trocha, Kristina Lefteri, Matthew Osborn, Julia Bals, Yannic C. Bartsch, Nathalie Bonheur, Timothy M. Caradonna, Josh Chevalier, Fatema Chowdhury, Thomas J. Diefenbach, Kevin Einkauf, Jon Fallon, Jared Feldman, Kelsey K. Finn, Pilar Garcia-Broncano, Ciputra Adijaya Hartana, Blake M. Paulina Kaplonek, Marshall Chenvang Jiang, Karpell. Hang Liu, Jinging Liu, Ngoc L. Ly, Ashlin Koscher, Xiaodong Lian, Michell, Yelizaveta Rassadkina, Kyra Seiger, Libera Sessa, Sally Shin, Nishant Singh, Weiwei Sun, Xiaoming Sun, Hannah J. Ticheli, Michael T. Waring, Alex L. Zhu, Galit Alter, Jonathan Z. Li, Daniel Lingwood, Aaron G. Schmidt, Mathias Lichterfeld, Bruce D. Walker, Xu G. Yu, Robert F. Padera Jr., Shiv Pillai, and the Massachusetts Consortium on Pathogen Readiness Specimen Working Group

Table S1 (Related to Figs. 1-4). Autopsied COVID-19 Patients studied by immunofluorescence and multispectral imaging

Duration of hospital stay	1-8 days	16-36 days	
	n=5	n=6	
Clinical Course			
Hospital stay duration (days) (mean ± s.d.)	3.6 (<u>+</u> 3.2)	25.8 (<u>+</u> 7.9)	
Demographics			
Age (years) (mean \pm s.d.)	66.2 (<u>+ 11.2)</u>	63.8 (<u>+</u> 9.3)	
Gender (% male)	60%	80%	
BMI (kg/m ²) (mean \pm s.d.)	33.0 (<u>+</u> 4.1)	30.2 (± 2.0)	
Maximum Laboratory Values			
$CRP (mg/L) (mean \pm s.d.)$	199.2 (<u>+</u> 102.3)	74.5 (<u>+</u> 44.8)	
LDH (U/L) (mean \pm s.d.)	590.0 (<u>+</u> 149.9)	423.0 (<u>+</u> 128.2)	
Ferritin (ug/L) (mean \pm s.d.)	703 (<u>+</u> 618.0)	410.8 (<u>+</u> 166.1)	
D-dimer (ng/mL) (range)	1374 to >4000	725 to >4000	

Body mass index (BMI), C-reactive protein (CRP), lactate dehydrogenase (LDH), standard deviation (s.d.)

Table S2 (Related to Figs. 1, 3 and 4). Non-COVID-19 Autopsied Controls

Age	Gender	Presumed Cause of Death		
77	F	Multi-organ system failure, pancreatitis		
78	M	Non-metastatic bladder malignancy		
64	M	Multi-organ system failure, thromboembolic infarcts		
78	M	Respiratory failure, treatment resistant MAC		
69	M	Cerebellar hemorrhage		
66	F	Fibrinous pericarditis, subpleural fibrosis		

Table S3 (Related to Figs. 2-4). Control Spleen Samples from Massachusetts General Hospital

Age	Gender	
61	M	
20	M	
54	F	
74	F	
59	M	
68	F	
58	M	

Table S4 (Related to Figs. 5-7). Summary of information on COVID-19 patients (n=68) in whom circulating B cells were analyzed by flow cytometry.

	Convalescence	Moderate	Severe (CRP int) CRP < 200 mg/L	Severe (CRP hi) CRP > 200 mg/L
	mean (<u>+</u> SD)	mean (<u>+</u> SD)	mean (<u>+</u> SD)	mean (<u>+</u> SD)
	n=39	n=4	n=10	n=15
Demographics				
Age (years)	44.8 (<u>+</u> 15.1)	35.5 (<u>+</u> 15.4)	59.3 (<u>+</u> 11.0)**	58.0 (<u>+</u> 13.9)**
Deceased (%)	0.0	0.0	10.0	13.3
Gender (% male)	28.2	50.0	60.0	66.7*
BMI (kg/m²)	24.5 (<u>+</u> 4.6)	31.1 (<u>+</u> 8.7)	35.7 (<u>+</u> 8.1)**	33.5 (<u>+</u> 8.7)**
Clinical Course				
Hospitalized due to COVID-19 (%)	0.0	0.0	100.0	100.0
Admitted to ICU due to COVID-19 (%)	0.0	0.0	70.0	86.7
Mechanically ventilated due to COVID-19 (%)	0.0	0.0	70.0	86.7
ECMO received due to COVID-19 (%)	0.0	0.0	0.0	6.7
Total Hospitalization (days)	na	na	26.5 (<u>+</u> 17.9)	55.5 (<u>+</u> 35.3) ⁺
Symptom Duration at Draw (days)	na	17.0 (<u>+</u> 9.7)	15.2 (<u>+</u> 12.3)	32.8 (<u>+</u> 14.4) ⁺⁺
Convalescence Period at Draw (days)	39.9 (<u>+</u> 15.0)	na	na	na
Maximum Laboratory Values				
CRP (mg/L)	na	na	103.3 (<u>+</u> 40.2)	314.6 (<u>+</u> 72.5)****
ESR (mm/h)	na	na	70.6 (<u>+</u> 44.1)	102.0 (<u>+</u> 36.7)
LDH (U/L)	na	na	526.5 (<u>+</u> 240.5)	707.2 (<u>+</u> 399.7)
Ferritin (ug/L)	na	na	2843.7 (<u>+</u> 5616.7)	3535.9 (<u>+</u> 3955.3)
D-dimer (ng/mL)	na	na	4399.0 (<u>+</u> 3157.5)	7669.3 (<u>+</u> 2691.4) ⁺
Day of Draw Laboratory Values				
ALC (K/uL)	1.50 (<u>+</u> 0.40)	na	1.40 (<u>+</u> 0.62)	1.55 (<u>+</u> 0.65)
CRP (mg/L)	na	na	39.8 (<u>+</u> 47.8)	74.4 (<u>+</u> 72.3)

Convalescence was defined as a clinically asymptomatic state on the date of blood draw, either from a baseline asymptomatic state or recuperated from moderate clinical symptoms of COVID-19. Moderate disease was defined as active clinical symptoms of COVID-19 on the date of blood draw that did not necessitate a hospital admission. Severe disease was defined as active clinical symptoms of COVID-19 on the date of blood draw that did necessitate a hospital admission. Severe disease was further subdivided by maximum CRP level during hospital admission as CRP < 200 mg/L 'CRP intermediate (int)' and CRP > 200 mg/L 'CRP high (hi).' Absolute lymphocyte count (ALC), Body mass index (BMI), C-reactive protein (CRP), extracorporeal membrane oxygenation (ECMO), erythrocyte sedimentation rate (ESR), lactate dehydrogenase (LDH), not available (na), standard deviation (SD), *significance by Student's t-test to 'convalescence' group, *significance by Student's t-test to 'severe (CRP int)' group.