Supplemental Table 3. ICU Day 1 plasma proteins that predict COVID19 outcome.

Num	Assay	Unipro ID	Function
1.	CXCL9	Q07325	A chemokine produced by airway epithelial cells in response to infection (1), also induced by IFNγ and in endothelial cells by TNFα (2). An agonist for CXCR3 on T cells and natural killer cells (2). Promotes NK, Th1, monocyte, DC, neutrophil, and eosinophil recruitment (1).
2.	ICOSLG	O75144	Inducible costimulator ligand is expressed by B cells, monocytes, DCs T cells and endothelial cells; TNFα is required for induction (3). It activates the inducible costimulator in the thymus and on activated T cells (3).
3.	CLM-1	Q8TDQ1	CMRF35-like molecule 1 is a receptor for phosphatidylserine presented on the outer membrane surface of apoptotic cells (4), that promotes macrophage and inhibits DC efferocytosis (5).
4.	IL12RB1	P42701	One of the two subunits that compose the IL-12 receptor (6), its signaling pathway activates STAT4 (7). Expressed primarily on activated T cells and NK cells, less so on dendritic cells and some B-cells (6, 7). This subunit is also shared with the IL-23 receptor (8).
5.	CD83	Q01151	Expressed on B and T cells, monocytes, DCs, microglia and neutrophils, and has soluble and membrane-bound forms (9). Membrane-bound CD83 is essential for CD4 ⁺ T cell development and inhibiting autoimmunity, soluble CD83 induces regulatory mechanisms for tolerance (9).
6.	CA12	O43570	Carbonic anhydrase 12 is membrane-associated glycoprotein that catalyzes the reversible hydration of carbon dioxide (10). CA12 is up-regulated by hypoxia, at least in tumor environments (11) and CA activity is associated with sleep apnea-related hypoxemia (12).
7.	FLRT2	O43155	Fibronectin leucine rich transmembrane protein 2 was first discovered in a screen for extracellular matrix proteins (13) and participates in homotypic cell-cell adhesion and with fibroblast growth factor receptor (14).
8.	ROR1	Q01973	A transmembrane receptor tyrosine kinase that is activated by Wnt family ligands and is mainly thought to be involved in organ/tissue genesis during development (15). Though recent evidence suggests it may be involved in proinflammatory p65 activation, at least in cancer (16).
9.	IL32	P24001	Expressed by PBMCs, epithelial cells and NKs, it upregulates other pro-inflammatory cytokines and has several isoforms (17). Airway epithelial cell production of IL32 is increased by viral infections and oxidative stress (1).
10.	NCS1	P62166	Neuronal calcium sensor 1 is a cytosolic protein involved in several cellular functions through binding partners and

			intracellular Ca ²⁺ regulation (18). It is highly expressed in neurons, but is not neuron-specific (18).
11.	S100A11	P31949	A cytosolic calcium-binding protein of the S100 family, it is involved in growth arrest in contact inhibition (19). It is expressed in a wide variety of cells and is secreted by an unconventional pathway (19). It is involved in cell-cell contacts and can promote cell migration in response to hypoxia-induced mitogenic factor (20).
12.	ANGPTL7	O43827	Angiopoietin-like protein 7 is an orphan ligand, but appears to be involved in hematopoietic stem cell regulation and self-renewal (21). Its serum concentration is higher in obese subjects compared to non-obese controls and can be lowered with exercise (22).
13.	CLMP	Q9H6B4	Coxsackievirus and adenovirus receptor-like membrane protein is a transmembrane glycoprotein involved in homophilic cell-cell adhesion and is expressed in a wide variety of tissues (23).
14.	IGF1R	P08069	A tyrosine kinase receptor expressed on T and B cells, macrophages, NK cells and granulocytes where its ligands, insulin-like growth factor 1 and 2, causes various effects such as proliferation, cytokine production and priming/activation (2).
15.	TOP2B	Q02880	DNA topoisomerase II beta is expressed in a wide variety of tissues and throughout the cell cycle (24). Mostly found in the cell's nucleus, it is one of the enzymes that catalyzes topological changes in DNA (24).
16.	FAM3B	P58499	Also called pancreatic derived factor (PANDER), it is highly expressed in pancreatic islets and high serum levels are associated with the progression of metabolic syndrome and type 2 diabetes (25).
17.	IL10.1	P22301	An important anti-inflammatory cytokine, expressed in virtually all immune cells except plasmacytoid DCs, to limit immune responses and prevent host damage (26).
18.	IL10	P22301	See "IL10.1" above.
19.	THY 1	P04216	Thymocyte differentiation antigen 1 is a glycoprotein expressed on the outer surface of many cell types including fibroblasts, T cells and activated endothelial cells and has a soluble form (27). Its function is cell and tissue-dependent, but is pro-fibrotic in pulmonary fibroblasts in pulmonary fibrosis (27).
20.	PVRL4	Q96NY8	Poliovirus receptor-related protein 4 also called nectin-4 is a cell-cell adhesion molecule in adherens junctions, overexpressed in several cancers (28).
21.	OPTC	Q9UBM4	Opticin is an extracellular matrix protein associated with collagen in the vitreous humor where it binds heparan and

	chondroitin sulfate (29, 30). It is an anti-angiogenic factor
	in retinas (30).

CXCR – CXC receptor

DC – dendritic cell

IFNγ – interferon gamma

IL – interleukin

NK – natural killer cell

PBMC – peripheral blood monocyte cell

STAT – signal transducer and activator of transcription

Th# – type # T helper cell

TNFα – tumor necrosis factor alpha

References

- Foti M, Locati M: Cytokine Effector Functions in Tissues [Internet]. San Diego, UNITED STATES: Elsevier Science & Technology; 2017. [cited 2020 Jun 9] Available from: http://ebookcentral.proquest.com/lib/west/detail.action?docID=4917635
- 2. Thomson AW, Lotze MT: The cytokine handbook. Amsterdam; Boston: Academic Press; 2003.
- 3. Richter G, Burdach S: ICOS: A New Costimulatory Ligand/Receptor Pair and Its Role in T-Cell Activion. *ORT* 2004; 27:91–95
- 4. Choi S-C, Simhadri VR, Tian L, et al.: Cutting Edge: Mouse CD300f (CLM-1) recognizes outer-membrane exposed phosphatidylserine and can promote phagocytosis. *J Immunol* 2011; 187:3483–3487
- 5. Tian L, Choi S-C, Lee H-N, et al.: Enhanced efferocytosis by dendritic cells underlies memory T-cell expansion and susceptibility to autoimmune disease in CD300f-deficient mice. *Cell Death Differ* 2016; 23:1086–1096
- 6. Presky DH, Yang H, Minetti LJ, et al.: A functional interleukin 12 receptor complex is composed of two β-type cytokine receptor subunits. *PNAS* 1996; 93:14002–14007

- 7. Trinchieri G: Interleukin-12 and the regulation of innate resistance and adaptive immunity. *Nature Reviews Immunology* 2003; 3:133–146
- 8. Wojno EDT, Hunter CA, Stumhofer JS: The immunobiology of the Interleukin-12 family: Room for discovery. *Immunity* 2019; 50:851–870
- 9. Grosche L, Knippertz I, König C, et al.: The CD83 Molecule An Important Immune Checkpoint [Internet]. *Front Immunol* 2020; 11[cited 2020 Jun 10] Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC7181454/
- 10. Waheed A, Sly WS, Doisy EA: Carbonic Anhydrase XII Functions in Health and Disease. *Gene* 2017; 623:33–40
- 11. Chiche J, Ilc K, Laferrière J, et al.: Hypoxia-Inducible Carbonic Anhydrase IX and XII Promote Tumor Cell Growth by Counteracting Acidosis through the Regulation of the Intracellular pH. *Cancer Res* 2009; 69:358–368
- 12. Wang T, Eskandari D, Zou D, et al.: Increased Carbonic Anhydrase Activity is Associated with Sleep Apnea Severity and Related Hypoxemia. *Sleep* 2015; 38:1067–1073
- 13. Lacy SE, Bönnemann CG, Buzney EA, et al.: Identification of FLRT1, FLRT2, and FLRT3: A Novel Family of Transmembrane Leucine-Rich Repeat Proteins. *Genomics* 1999; 62:417–426
- 14. Karaulanov EE, Böttcher RT, Niehrs C: A role for fibronectin-leucine-rich transmembrane cell-surface proteins in homotypic cell adhesion. *EMBO Rep* 2006; 7:283–290
- 15. Stricker S, Rauschenberger V, Schambony A: Chapter Four ROR-Family Receptor Tyrosine Kinases [Internet]. In: Jenny A, editor(s). Current Topics in Developmental Biology. Academic Press; 2017. p. 105–142.[cited 2020 Jun 10] Available from: http://www.sciencedirect.com/science/article/pii/S007021531630179X
- 16. Lopez-Bergami P, Barbero G: The emerging role of Wnt5a in the promotion of a proinflammatory and immunosuppressive tumor microenvironment. *Cancer Metastasis Rev* 2020;
- 17. Catalan-Dibene J, McIntyre LL, Zlotnik A: Interleukin 30 to Interleukin 40. *Journal of Interferon & Cytokine Research* 2018; 38:423
- 18. Boeckel GR, Ehrlich BE: NCS-1 is a Regulator of Calcium Signaling in Health and Disease. *Biochim Biophys Acta Mol Cell Res* 2018; 1865:1660–1667
- 19. Sakaguchi M, Huh N: S100A11, a dual growth regulator of epidermal keratinocytes. *Amino Acids* 2011; 41:797–807
- 20. Gross SR, Sin CGT, Barraclough R, et al.: Joining S100 proteins and migration: for better or for worse, in sickness and in health. *Cell Mol Life Sci* 2014; 71:1551–1579

- 21. Kadomatsu T, Oike Y: Roles of angiopoietin-like proteins in regulation of stem cell activity. *J Biochem* 2019; 165:309–315
- 22. Abu-Farha M, Cherian P, Al-Khairi I, et al.: Plasma and adipose tissue level of angiopoietin-like 7 (ANGPTL7) are increased in obesity and reduced after physical exercise. *PLOS ONE* 2017; 12:e0173024
- 23. Matthäus C, Langhorst H, Schütz L, et al.: Cell-cell communication mediated by the CAR subgroup of immunoglobulin cell adhesion molecules in health and disease. *Molecular and Cellular Neuroscience* 2017; 81:32–40
- 24. Austin CA, Lee KC, Swan RL, et al.: TOP2B: The First Thirty Years [Internet]. *Int J Mol Sci* 2018; 19[cited 2020 Jun 11] Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6163646/
- 25. Zhang X, Yang W, Wang J, et al.: FAM3 gene family: A promising therapeutical target for NAFLD and type 2 diabetes. *Metabolism* 2018; 81:71–82
- 26. Howes A, Stimpson P, Redford P, et al.: Interleukin-10: Cytokines in Anti-inflammation and Tolerance [Internet]. In: Yoshimoto T, Yoshimoto T, editor(s). Cytokine Frontiers: Regulation of Immune Responses in Health and Disease. Tokyo: Springer Japan; 2014. p. 327–352.[cited 2020 Jun 9] Available from: https://doi.org/10.1007/978-4-431-54442-5_13
- 27. Bradley JA, Ramirez G, Hagood JS: Roles and Regulation of Thy-1, a Context-Dependent Modulator of Cell Phenotype. *Biofactors* 2009; 35:258–265
- 28. Takai Y, Miyoshi J, Ikeda W, et al.: Nectins and nectin-like molecules: roles in contact inhibition of cell movement and proliferation. *Nat Rev Mol Cell Biol* 2008; 9:603–615
- 29. Le Goff MM, Bishop PN: Focus on Molecules: Opticin. *Experimental Eye Research* 2007; 85:303–304
- 30. Bishop PN: The role of extracellular matrix in retinal vascular development and preretinal neovascularization. *Experimental Eye Research* 2015; 133:30–36