## **Immunologists struggling to understand COVID-19**

## ARE IN IT FOR THE LONG HAUL

Researchers and clinicians from fifteen institutions across the country are working together to enroll nearly 2000 COVID-19 patients in a clinical trial examining acute and long-term immune responses to the virus. The goal of the study, dubbed Immunotyping Assessment in a COVID-19 Cohort (IMPACC), is to characterize immune signatures to differentiate infection severity and explore long-term effects. Understanding immune response heterogeneity in COVID-19 patients could improve treatment and help "long-haulers" combat lingering effects.

IMPACC
researchers and
clinicians hope to help
clinicians predict which
patients may have the
most severe infections,
what treatment is best,
and when the
treatment should
be administered.

IMPACC
is part of the
Human Immunology
Project Consortium (HIPC)
established by the NIAID. The
HIPC will develop a publicly
available resource detailing
varied immune responses to
infections, treatments,
and vaccines.

Patients
with severe
COVID-19 infection
have an intense extrafollicular B-cell response
akin to that found in autoimmune disorders such
as systemic lupus
erythematosus.<sup>1,2</sup>





Clinicians collect nasal swabs and blood samples from patients throughout their hospitalization and at follow-up visits once every three months. They also collect lower airway secretions from patients using ventilators.

COVID-19 infected patients develop autoantibodies against the following:

- Phospholipids and carbohydrates<sup>3, 4, 5</sup>
- · Rheumatoid factor<sup>3</sup>
- Blood plasma proteins important for blood clotting<sup>3,6</sup>
- Cytokines, chemokines, and complement components<sup>7,8</sup>

Researchers will collect cells from individual patients to quantify the population of immune cells and immune related proteins such as cytokines and interferons in the patient samples at the collected time points. The researchers will use several approaches, including cytometry by time of flight, next generation sequencing, and proteomics to analyze the samples.

## REFERENCES

- Woodruff, M.C. et al. Extrafollicular B cell responses correlate with neutralizing antibodies and morbidity in COVID-19. Nat Immuno. 21: 1506–1516. (2020).
- Woodruff, M.C. et al. Clinically identifiable autoreactivity is common in severe SARS-CoV-2 Infection. medRxiv. (2020).
- Butler, D.L. et al. Abnormal antibodies to self-carbohydrates in SARS-CoV-2 infected patients. bioRxiv. (2020)

- Zuo, Y. et al. Prothrombotic autoantibodies in serum from patients hospitalized with COVID-19. Sci Transl Med. 12 (570). (2020).
- Wang, E.Y. et al. Diverse Functional Autoantibodies in Patients with COVID-19. medRxiv. (2020).
- Bastard, P. et al. Autoantibodies against type I IFNs in patients with life-threatening COVID-19. Science 370 (6515). (2020).