

**Supplemental Table 1: Summary of infectious, hematologic, hereditary, metabolic, and autoimmune work-ups (with course of pharmacologic agents included)**

| Infectious Work-Up                                      |                                                                                            |
|---------------------------------------------------------|--------------------------------------------------------------------------------------------|
| Study                                                   | Result                                                                                     |
| Clostridium difficile DNA, feces                        | tcdB gene DNA detected                                                                     |
| SARS-CoV-2 PCR                                          | Not detected                                                                               |
| Stool culture                                           | No salmonella, shigella, campylobacter, E coli O157:H7                                     |
| Blood cultures (multiple)                               | No growth                                                                                  |
| Urine culture                                           | Escherichia coli (10K-15K cfu/mL; pansusceptible)                                          |
| Intestinal Ova & Parasites                              | No ova or parasites found                                                                  |
| Enteric parasite PCR panel                              | No E. histolytica, Giardia, Cryptosporidium PCR detected                                   |
| Chlamydia + Gonococcus DNA, urine                       | Negative                                                                                   |
| HIV 1/0/2 antibody, serum                               | Negative                                                                                   |
| Viral Hepatitis panel                                   | Negative for HBV S Ag, HBV C Ab, HAV Ab IgM, HCV Ab                                        |
| Quantiferon tuberculosis Gold                           | Negative                                                                                   |
| Hantavirus antibody IgG/IgM, serum                      | Negative                                                                                   |
| Leptospira species antibody IgM, serum                  | Negative                                                                                   |
| Coxiella burnetii Phase 1 & 2 antibody (IgM/IgG), serum | All titers <1:16                                                                           |
| Meningitis/Encephalitis panel, CSF                      | Negative for all targets                                                                   |
| Cryptococcus species antigen, CSF                       | Negative                                                                                   |
| VDRL antibody, CSF                                      | Non-reactive                                                                               |
| West Nile Virus antibody IgG/IgM, CSF                   | Negative                                                                                   |
| Arbovirus panel, CSF                                    | Negative for California Encephalitis, East Equine, St Louis, West Equine, West Nile        |
| Coccidioides immitis antibody, IgG/IgM, CSF             | Negative                                                                                   |
| Rickettsia Panel (Typhus, Rickettsiae), serum           | Titers <1:64                                                                               |
| Syphilis Screen                                         | Non-reactive for Treponema pallidum antibody                                               |
| Cryptococcus species antigen, CSF                       | Negative                                                                                   |
| West Nile Virus antibody IgG/IgM, Serum                 | Negative                                                                                   |
| Dengue Virus antibody IgG/IgM, Serum                    | Negative                                                                                   |
| Histoplasma capsulatum antigen                          | Negative                                                                                   |
| Legionella pneumophila antigen, Urine                   | Negative                                                                                   |
| SARS-CoV-2 antibody, IgM                                | Negative                                                                                   |
| SARS-CoV02 antibody, IgA                                | Negative                                                                                   |
| Hematologic Work-Up                                     |                                                                                            |
| Direct Antiglobulin Test                                | Negative                                                                                   |
| D-dimer                                                 | >20.00 mcg/mL                                                                              |
| Peripheral smear                                        | Marked schistocytes                                                                        |
| Haptoglobin (nadir)                                     | <15 mg/dL                                                                                  |
| Lactate dehydrogenase (peak value)                      | 1,984 U/L (reference range: 135-225 U/L)                                                   |
| Fibrinogen (nadir)                                      | 78 mg/dL                                                                                   |
| Platelet count (nadir)                                  | 42 x10 <sup>3</sup>                                                                        |
| Coagulation panel (peak values)                         | Protime 19.5 seconds, APTT 54.0 seconds, INR 1.6                                           |
| Porphyrins Quantitative Analysis, Urine                 | Within normal limits                                                                       |
| Porphobilinogen, Urine                                  | 1.1 mg/dL (reference range: 0.0-2.0 mg/dL)                                                 |
| ADAMTS13 Activity                                       | 61.9% (reference range: >66.8%)                                                            |
| Immunoglobulins Quantitative Analysis, serum            | IgG 410 mg/dL (reference range: 700-1600 mg/dL), IgA/IgM within normal limits              |
| Peripheral blood flow cytometry                         | 6% monocytes with dim CD56 expression, no monoclonal B-cell or T-cell population, no CD34+ |
| Tissue Exam (colon)                                     | Pseudomembranous colitis with superficial mucosal thrombotic microangiopathy; no carcinoma |
| Hereditary & Metabolic Work-up                          |                                                                                            |
| Ceruloplasmin                                           | 17 mg/dL (reference range: 16-45 mg/dL)                                                    |
| Creatine kinase                                         | 829 U/L (reference range: 26-192 U/L)                                                      |
| Ammonia, serum                                          | 26 mcmol/L (reference range: 11-51 mcmol/L)                                                |
| Amino Acid panel                                        | Within normal limits                                                                       |
| Pyruvate, serum                                         | 0.6 mg/dL (reference range: 0.3-0.7 mg/dL)                                                 |

|                                                    |                                                                                             |
|----------------------------------------------------|---------------------------------------------------------------------------------------------|
| Acylcarnitine Profile Analysis, serum              | Non-specific pattern                                                                        |
| Organic Acids Profile Analysis, Urine              | Non-specific pattern                                                                        |
| Fulgent Diagnostics (Genome Sequencing)            | No clinically significant abnormalities (including C3, CAPG, CD46, CFB, CFH, CFI, and THBD) |
| <b>Autoimmune &amp; Immunologic Work-Up</b>        |                                                                                             |
| Autoimmune Encephalitis panel                      | Negative for all 18 targets tested                                                          |
| Complement C3                                      | 57 mg/dL (reference range: 90-180 mg/dL)                                                    |
| Complement C4                                      | <12 mg/dL (reference range: 10-40 mg/dL)                                                    |
| Complement CH50                                    | 40 U/mL (reference range: >41 U/mL)                                                         |
| Factor H Autoantibody (FHAB)                       | 2.5% (reference range: 0.0-7.3%)                                                            |
| CNS Demyelinating Disease Evaluation, serum        | Negative for NMO, AQP4, and MOG                                                             |
| Thyroid-stimulating Hormone (TSH)                  | 7.2 mIU/mL (reference range: 0.27-5.00 mIU/mL)                                              |
| Thyroxine Free (FT4)                               | 1.05 ng/dL (reference range: 0.6-1.8 ng/dL)                                                 |
| Thyroglobulin antibody                             | 1.76 IU/mL (reference range: 0-4.1 IU/mL)                                                   |
| Thyroperoxidase antibody                           | 1.18 IU/mL (reference range: 0-5.6 IU/mL)                                                   |
| Cryoglobulin, serum                                | None detected                                                                               |
| Antinuclear Antibody (ANA) Reflex Cascade          | Negative                                                                                    |
| Smooth muscle antibody                             | 1:20 titer                                                                                  |
| Antimitochondrial antibody                         | Non-reactive                                                                                |
| Vasculitis panel                                   | Negative for Myeloperoxidase, Proteinase 3, and Glomerular basement membrane antibodies     |
| <b>Pharmacologic Agents During Hospital Course</b> |                                                                                             |
| <b>Medication (Route, Dose)</b>                    | <b>Course (Listed by Hospital Day)</b>                                                      |
| Acetaminophen (IV, 1000 mg)                        | 1 – 3                                                                                       |
| Ampicillin (IV, 2gm)                               | 7                                                                                           |
| Bromocriptine (PO, 2.5 mg)                         | 7 – 10                                                                                      |
| Ceftriaxone (IV, 1-2 gm)                           | 1 – 3, 7                                                                                    |
| Cyproheptadine (PO, 12 mg)                         | 7                                                                                           |
| Dexamethasone (IV, 4-10 mg)                        | 5 – 12                                                                                      |
| Dexmetomidine (IV, titrated)                       | 4 – 7                                                                                       |
| Doxycycline (IV, 100 mg)                           | 7 – 30                                                                                      |
| Eculizumab (IV, 900 mg)                            | 13, 20, 27                                                                                  |
| Enoxaparin (SQ, 40 mg)                             | 1 – 3, 29 – 33                                                                              |
| Fentanyl (IV, titrated)                            | 7 – 12                                                                                      |
| Folic acid (PO, 1 mg)                              | 18 – 33                                                                                     |
| Gabapentin (PO, 100 mg)                            | 2                                                                                           |
| Haloperidol (IV, 2-5 mg)                           | 5 – 6                                                                                       |
| Heparin (SQ, 5000 units)                           | 4 – 29                                                                                      |
| Hydromorphone (IV, 0.2 mg)                         | 4 – 5                                                                                       |
| Lacosamide (IV, 200 mg)                            | 7 – 9                                                                                       |
| Levetiracetam (IV, 1000 mg)                        | 7 – 30                                                                                      |
| Lorazepam (IV, 5 mg)                               | 7 – 8                                                                                       |
| Methylprednisolone (IV, 1 gm)                      | 12 – 17                                                                                     |
| Metronidazole (IV, 500 mg)                         | 2 – 11                                                                                      |
| Morphine (IV, 2-4 mg)                              | 2 – 9                                                                                       |
| Mupirocin (intranasal, 2%)                         | 3 – 9                                                                                       |
| Olanzapine (PO, 5 mg)                              | 6 – 7                                                                                       |
| Ondansetron (IV, 4 mg)                             | 1, 3, 5                                                                                     |
| Oxycodone (PO, 5-10 mg)                            | 2 – 6, 13 – 15                                                                              |
| Pantoprazole (IV, 40 mg)                           | 4 – 19                                                                                      |
| Prednisone (PO, 20-60)                             | 17 – 33                                                                                     |
| Promethazine (IV, 25 mg)                           | 1 – 3                                                                                       |
| Propofol (IV, titrated)                            | 7 – 12                                                                                      |
| Ramelteon (PO, 8 mg)                               | 5 – 7                                                                                       |
| Thiamine (IV, 250-500 mg)                          | 4 – 11                                                                                      |
| Vancomycin (PO, 125-500 mg)                        | 1 – 16                                                                                      |
| Vancomycin (IV, 1.25 gm)                           | 7                                                                                           |