

CASE STUDIES

Infection-triggered autoimmune encephalopathy syndromes

Volume 1

The following case studies demonstrate the clinical utility of the Autoimmune BrainPanel* as an aid in identifying an underlying autoimmune etiology in patients with certain neuropsychiatric symptoms. Test results can support a clinician's diagnosis with laboratory evidence.



Case Study #1 - Autoimmune-induced neuropsychiatric symptoms in an 8-year-old female

Antibiotic treatment leads to resolution of multiple neurologic and psychiatric symptoms

READ

Case Study #2 - Gradual onset of tics and severe cognitive regression in 20-year-old male

Initially diagnosed with bipolar disorder, testing reveals immune-mediated condition

READ

Case Study #3 - Severe OCD and tics improve with immune-modulatory treatment

Clinical and laboratory results support autoimmunebased illness

READ

Case Study #4 - 20-year-old female with OCD improves with rituximab treatment

Testing revealed elevated autoantibodies against Dopamine D2L receptor

READ

Case Study #5 - Multiple neuropsychiatric symptoms improve with immune-modulatory treatment

Reduction in symptoms following treatment with anti-inflammatory drugs and IVIg therapy

READ

The Autoimmune Brain Panel™ - An Overview

READ

Autoimmune-induced neuropsychiatric symptoms in an 8-year-old female

Antibiotic treatment leads to resolution of multiple neurologic and psychiatric symptoms

Presentation: 8-year-old girl with abrupt onset of obsessive-compulsive behaviors, vocal tics, poor concentration, sensory/motor abnormalities, emotional lability, regression, aggression, and dysgraphia. Parents report urinary and sleep problems.

Course of Illness: Relapsing and remitting symptoms with improvement after 2-3 days of azithromycin.

Previous Diagnoses: Anxiety, OCD, unspecified disorder of immune mechanism

Previous Lab Results: ASO <50, Anti-DNase B <50, normal IgG titers

Medical History: Significant for two recent streptococcal infections with high fever, sore throat, sinus pain and cough. History of frequent ear infections, sinus infections and idiopathic fevers.

Family History: Positive for mycoplasma infections, autoimmune disorders, allergies, frequent bacterial and viral infections.

Pre-Treatment Autoimmune Brain Panel™ Results: Elevated anti-Lysoganglioside GM1 autoantibodies 640 (normal range 80-320), elevated anti-Tubulin autoantibodies 2,000 (normal range 250-1,000), and borderline anti-Dopamine Receptor D1 autoantibodies 2,000 (normal range 500-2,000)

Treatment: Azithromycin with rapid symptom improvement.

Post-Treatment Autoimmune Brain Panel™ Results:

Anti-Lysoganglioside GM1 returned to normal (147); anti-Tubulin returned to normal (250) and anti-Dopamine Receptor D1 autoantibodies returned to normal (500)

Status: All symptoms resolved. Returned to baseline after 2-3 days of treatment.



Autoimmune Brain Panel[™] (formerly known as the Cunningham Panel[™])

PRE-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor
Autoantibodies

Borderline 2000

Anti-Dopamine D2L Receptor Autoantibodies

Normal 2000

Anti-Lysoganglioside GM1 Autoantibodies

Elevated 640

Anti-Tubulin Autoantibodies

Elevated 2000

CaMKinase II

Normal 95

POST-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor	Normal
Autoantibodies	500
Anti-Dopamine D2L Receptor	Normal
Autoantibodies	2000
Anti-Lysoganglioside	Normal
GM1 Autoantibodies	40
Anti-Tubulin	Normal
Autoantibodies	250

CaMKinase II

Normal 112

An 8-year-old girl with a history of strep, sinus and ear infections presented with numerous neuropsychiatric symptoms exhibited clinical and laboratory response to antibiotics.

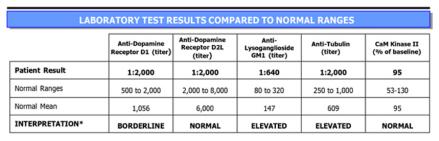


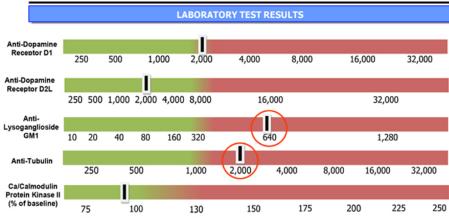
Autoimmune Brain Panel[™]test results (formerly known as the Cunningham Panel[™])

PRE-TREATMENT:

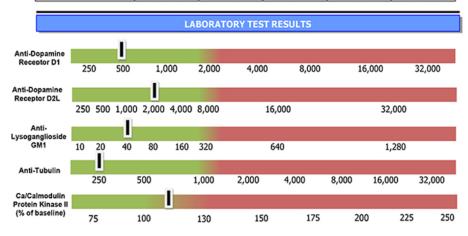
Patient Symptomatic

Patient symptoms pre-treatment consistent with elevated autoantibodies to Lysoganglioside GM1 (tics, dysgraphia), elevated anti-Tubulin autoantibodies 2000 (OCD, poor concentration), and borderline Dopamine D1 autoantibodies (anxiety, sleep problems, emotional lability, aggression, regression).





LABORATORY TEST RESULTS COMPARED TO NORMAL RANGES					
	Anti-Dopamine Receptor D1 (titer)	Anti-Dopamine Receptor D2L (titer)	Anti- Lysoganglioside GM1 (titer)	Anti-Tubulin (titer)	CaM Kinase II (% of baseline)
Patient Result	1:500	1:2,000	1:40	1:250	112
Normal Ranges	500 to 2,000	2,000 to 8,000	80 to 320	250 to 1,000	53-130
Normal Mean	1,056	6,000	147	609	95
INTERPRETATION*	NORMAL	NORMAL	NORMAL	NORMAL	ELEVATED



POST-TREATMENT:

Symptom Resolution

Symptom resolution complete. All Autoimmune Brain Panel™ markers within normal limits.



Gradual onset of tics and severe cognitive regression in 20-year-old male

Initially diagnosed with bipolar disorder, testing reveals immune-mediated condition

Presentation: 20-year-old male with gradual onset of tics, inability to concentrate, sensory abnormalities, emotional lability, severe cognitive regression, separation anxiety, developmental regression, sleep disturbances, handwriting disturbance and aggressiveness.

Previous Diagnoses: Unspecified bipolar disorders

Previous Lab Results: Positive for Lyme disease, Babesia, and Rickettsia

Medical History: Rocky Mountain Spotted Fever, multiple ear and strep infections throughout childhood, known immunodeficiency.

Family History: Positive for rheumatoid arthritis, asthma and eczema.

Pre-Treatment Autoimmune Brain Panel™ Results: Elevated anti-Dopamine Receptor D1 autoantibodies - 4,000 (normal range 500-2,000), borderline anti-Dopamine Receptor D2L - 8,000 (normal range 2,000-8,000), elevated anti-Tubulin autoantibodies - 2,000 (normal range 250-1,000), elevated Calcium/calmodulin-dependent protein kinase II (CaMKII) - 140 (normal range 53-130)

Treatment: Plasmapheresis followed by multiple IVIg treatments.

Post-Treatment Autoimmune Brain Panel™ Results: Anti-Dopamine Receptor D1 autoantibodies decreased to borderline (2,000), anti-Dopamine Receptor D2L autoantibodies remained at borderline (8,000), anti-Tubulin autoantibodies decreased to borderline (1,000), Calcium/calmodulin-dependent protein kinase II (CaMKII) decreased to borderline (124)

Status: Overall cognitive abilities improved, with mother reporting patient's math skills returned after many years.



Autoimmune Brain Panel™

(formerly known as the Cunningham Panel™)

PRE-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor
Autoantibodies

Elevated 4000

Anti-Dopamine D2L Receptor Autoantibodies

Borderline

8000

Anti-Lysoganglioside GM1 Autoantibodies Normal

160

Anti-Tubulin Autoantibodies Elevated 2000

CaMKinase II

Elevated 140

POST-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor
Autoantibodies

Borderline 2000

Anti-Dopamine D2L Receptor Autoantibodies

Borderline 8000

Anti-Lysoganglioside GM1 Autoantibodies

Normal 160

Anti-Tubulin Autoantibodies

Borderline 1000

CaMKinase II

Borderline 124

A 20-year-old male with gradual onset of tics, severe cognitive regression and multiple neuropsychiatric symptoms, who tested positive for multiple tick-borne illnesses, had elevated anti-Dopamine Receptor D1 and anti-Tubulin autoantibodies, along with an elevated cell stimulation assay (CaMKII) suggesting an infection-triggered autoimmune-based etiology.



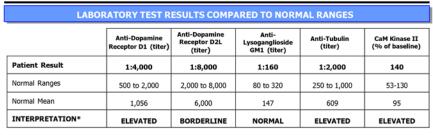
Autoimmune Brain Panel[™] test results (formerly known as the Cunningham Panel[™])

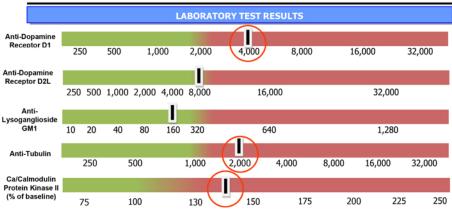
PRE-TREATMENT:

Patient Symptomatic

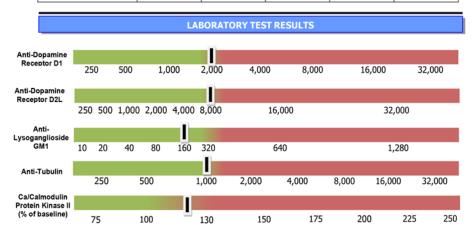
Elevated Dopamine Receptor D1 autoantibodies (4,000) consistent with emotional lability, anxiety, regression, sleep problems and aggressiveness; Tubulin autoantibodies (2,000) consistent with poor concentration and cognitive regression.

Calcium/calmodulin-dependent protein kinase II (CaMKII) at 140 correlates with tics, sensory abnormalities, and dysgraphia.





LABORATORY TEST RESULTS COMPARED TO NORMAL RANGES					
	Anti-Dopamine Receptor D1 (titer) Anti-Dopamine Receptor D2L (titer) Anti-Lysoganglioside GM1 (titer) Anti-Tubulin (titer) CaM Kinase II (% of baseline)				
Patient Result	1:2,000	1:8,000	1:160	1:1,000	124
Normal Ranges	500 to 2,000	2,000 to 8,000	80 to 320	250 to 1,000	53-130
Normal Mean	1,056	6,000	147	609	95
INTERPRETATION*	BORDERLINE	BORDERLINE	NORMAL	BORDERLINE	BORDERLINE



POST-TREATMENT:

Symptom Resolution

Borderline Dopamine
Receptor D1 autoantibodies
(2,000), Dopamine Receptor
D2L autoantibodies (8,000),
Tubulin autoantibodies (1,000)
and CaMKII (124) consistent
with resolution of symptoms.



Severe OCD and tics improve with immune-modulatory treatment

Clinical and laboratory results support autoimmune-based illness

Presentation: 16-year-old female with abrupt onset at age 11 of severe OCD (food restriction), tics and choreiform movements, poor concentration, significant cognitive and academic decline, sensory abnormalities, severe mood dysregulation, urinary urgency and frequency, and dysgraphia. Aggressive behaviors prompted 6 calls to local police for assistance.

Course of Illness: Relapsing and remitting symptoms with three major episodes reported. Sudden increase in symptoms with strep exposure.

Previous Diagnoses: Autoimmune disease NOS, post-infectious encephalitis and encephalomyelitis, other diseases involving the immune system NOS

Previous Lab Results: Positive Lyme Western blot, elevated Mycoplasma and Coxsackie titers

Medical History: Non-contributory

Family History: Maternal and paternal grandmothers with autoimmune disorders, history of rheumatic fever. Sibling with Pediatric Autoimmune Neuropsychiatric Disorders Associated with Strep infection (PANDAS).

Pre-Treatment Autoimmune Brain Panel™ Results: Elevated Calcium/calmodulin-dependent protein kinase II (CaMKII) 167 (normal range 53-130)

Treatment: Plasmapheresis and IVIg with symptom improvement, low dose Zoloft, Abilify for maintenance.

Post-Treatment Autoimmune Brain Panel™ Results: Calcium / calmodulin-dependent protein kinase II (CaMKII) returned to borderline (123)

Status: Symptom improvement.



Autoimmune Brain Panel™

(formerly known as the Cunningham Panel™)

PRE-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor Borderline 2000 Autoantibodies Anti-Dopamine D2L Receptor Normal 4000 **Autoantibodies** Anti-Lysoganglioside Normal **GM1** Autoantibodies 40 Anti-Tubulin Borderline 1000 **Autoantibodies** Elevated CaMKinase II 167

POST-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor	Normal
Autoantibodies	1000
Anti-Dopamine D2L Receptor	Normal
Autoantibodies	4000
Anti-Lysoganglioside	Normal
GM1 Autoantibodies	80
Anti-Tubulin	Borderline
Autoantibodies	1000
CaMKinase II	Borderline 123

A 16-year-old female with severe food restriction, OCD, tics, choreiform movements, poor concentration, cognitive and academic decline, sensory abnormalities, mood dysregulation, urinary urgency and frequency, and dysgraphia exhibited clinical and laboratory response to plasmapheresis and IVIg.

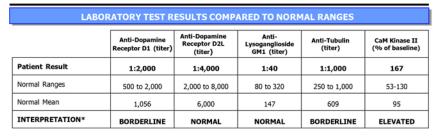


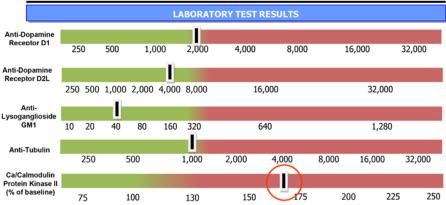
Autoimmune Brain Panel[™]test results (formerly known as the Cunningham Panel™)

PRE-TREATMENT:

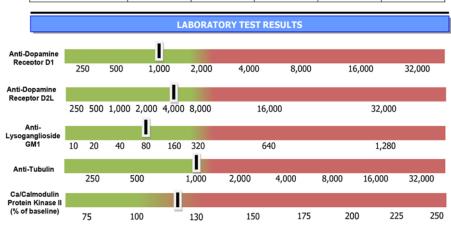
Patient Symptomatic

Patient symptoms likely due to CaMKII activation and subsequent increase in dopamine, epinephrine, and norepinephrine neurotransmitters resulting in adrenergic stimulation. Elevation in CaMKII associated with likely active infection.





LABORATORY TEST RESULTS COMPARED TO NORMAL RANGES					
	Anti-Dopamine Receptor D1 (titer) Anti-Dopamine Receptor D2L (titer) Anti-Dopamine Receptor D2L Lysoganglioside GM1 (titer) Anti-Tubulin (% of baseline)				
Patient Result	1:1,000	1:4,000	1:80	1:1,000	123
Normal Ranges	500 to 2,000	2,000 to 8,000	80 to 320	250 to 1,000	53-130
Normal Mean	1,056	6,000	147	609	95
INTERPRETATION*	NORMAL	NORMAL	NORMAL	BORDERLINE	BORDERLINE



POST-TREATMENT:

Symptom Resolution

Symptom resolution after multiple plasma exchanges to mechanically remove antibodies followed by IVIg to replenish healthy antibodies.



20-year-old female with OCD improves with rituximab treatment

Testing revealed elevated autoantibodies against Dopamine D2L receptor

Presentation: 20-year-old female with gradual onset of OCD, poor concentration, sensory and motor abnormalities, emotional lability, separation anxiety, developmental regression, urinary frequency and urgency, sleep disturbances, and aggression.

Course of Illness: Symptoms relapsing and remitting with very frequent flares.

Previous Diagnoses: Idiopathic peripheral neuropathy, reflex sympathetic dystrophy, variants of migraine, Lyme disease, disorder of autonomic nervous system

Previous Lab Results: Positive for strep, Lyme disease, Babesia, Mycoplasma, Epstein-Barr virus, Herpes Zoster virus

Medical History: Frequent strep infections since childhood, low quantitative immunoglobulins, and dental infections.

Family History: Positive for autoimmune disease and allergies.

Pre-Treatment Autoimmune Brain Panel™ Results: Elevated anti-dopamine receptor D2L autoantibodies 16,000 (normal range 2,000-8,000), elevated Calcium/calmodulin-dependent protein kinase II (CaMKII) 144 (normal range 53-130)

Treatment: Rituximab (4 courses) and ozone treatments with significant symptom reduction.

Post-Treatment Autoimmune Brain Panel™ Results: Anti-dopamine receptor D2L autoantibodies returned to normal (500), Calcium / calmodulin-dependent protein kinase II (CaMKII) levels returned to normal (111)

Status: Significant reduction in symptoms.



Autoimmune Brain Panel[™] (formerly known as the Cunningham Panel[™])

PRE-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor Normal 1000 Autoantibodies Anti-Dopamine D2L Receptor Elevated **Autoantibodies** 16000 Anti-Lysoganglioside Normal **GM1** Autoantibodies 80 Anti-Tubulin Normal 500 **Autoantibodies** Elevated CaMKinase II

POST-TREATMENT TEST RESULTS

144

Anti-Dopamine D1 Receptor	Normal
Autoantibodies	1000
Anti-Dopamine D2L Receptor	Normal
Autoantibodies	500
Anti-Lysoganglioside	Normal
GM1 Autoantibodies	40
Anti-Tubulin	Normal
Autoantibodies	250
CaMKinase II	Normal 111

A 20-year-old female presented with gradual onset OCD and multiple neuropsychiatric symptoms. Testing revealed elevated autoantibodies directed against the Dopamine D2L receptor and elevated CaMKII, suggesting an autoimmune component. Elimination of memory B cells with rituximab resulted in a significant reduction in symptoms.

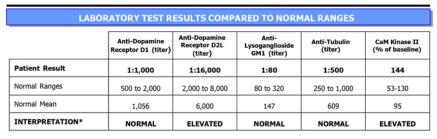


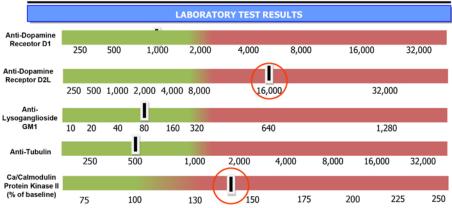
Autoimmune Brain Panel[™] test results (formerly known as the Cunningham Panel[™])

PRE-TREATMENT:

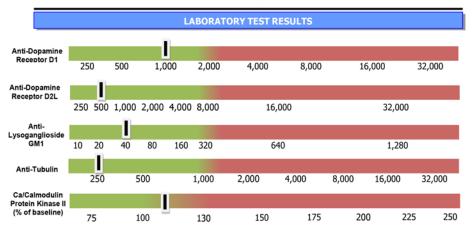
Patient Symptomatic

Patient symptomatic likely due to autoantibodies directed against D2L receptors resulting in motor abnormalities and increased CaMKII activation associated with OCD, poor concentration, emotional lability, separation anxiety, developmental regression, urinary issues, sleep disturbances, and aggression.





LABORATORY TEST RESULTS COMPARED TO NORMAL RANGES					
	Anti-Dopamine Receptor D2L Lysoganglioside (titer) GM1 (titer) Anti-Tubulin (titer) CaM Kinase II (% of baseline)				
Patient Result	1:1,000	1:500	1:40	1:250	111
Normal Ranges	500 to 2,000	2,000 to 8,000	80 to 320	250 to 1,000	53-130
Normal Mean	1,056	6,000	147	609	95
INTERPRETATION*	NORMAL	NORMAL	NORMAL	NORMAL	NORMAL



POST-TREATMENT:

Symptom Resolution

Symptom resolution after four courses of Rituxan (rituximab), a chemotherapeutic monoclonal antibody targeting and eliminating memory B cells.



Multiple neuropsychiatric symptoms improve with immune-modulatory treatment

Reduction in symptoms following treatment with anti-inflammatory drugs and IVIg therapy

Presentation: 17-year-old male with abrupt onset of OCD, inability to concentrate, sensory and motor abnormalities, emotional lability, separation anxiety, developmental regression, sleep disturbances, handwriting disturbance, aggressiveness, and choreiform movements.

Course of Illness: Relapsing and remitting symptoms.

Previous Diagnoses: Panic disorder, emotional disorders with onset specific to childhood, selective deficiency of immunoglobulin A (IgA), personal history of traumatic brain injury, other encephalitis and encephalomyelitis

Previous Lab Results: High Coxsackie, Parvovirus and HHV6 titers, hypogammaglobinemia, MARCONS positive

Medical History: Chronic sinusitis and frequent viral infections.

Family History: Positive for rheumatic fever, tics, OCD, autoimmune disorders, hyper and hypothyroidism and allergies.

Pre-Treatment Autoimmune Brain Panel™ Results: Borderline anti-Tubulin autoantibodies - 1,000 (normal range 250-1,000), and elevated Calcium/calmodulin-dependent protein kinase II (CaMKII) - 184 (normal range 53-130)

Treatment: NSAIDs and multiple IVig infusions.

Post-Treatment Autoimmune Brain Panel™ Results: Anti-Tubulin autoantibodies remained borderline (1,000) and Calcium / calmodulin-dependent protein kinase II (CaMKII) returned to normal (113)

Status: Symptom improvement with anti-inflammatory drugs and IVIG treatment.



Autoimmune Brain Panel™

(formerly known as the Cunningham Panel™)

PRE-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor Autoantibodies

Normal 500

Anti-Dopamine D2L Receptor Autoantibodies

Normal 4000

Anti-Lysoganglioside GM1 Autoantibodies

Normal 80

Anti-Tubulin Autoantibodies

Borderline 1000

CaMKinase II

Elevated 184

POST-TREATMENT TEST RESULTS

Anti-Dopamine D1 Receptor
Autoantibodies

Anti-Dopamine D2L Receptor
Autoantibodies

Anti-Lysoganglioside
GM1 Autoantibodies

Anti-Tubulin

Borderline
Borderline
Borderline

Autoantibodies

1000

CaMKinase II

Normal 113

A 17-year-old male with abrupt onset OCD and other neuropsychiatric symptoms and a history of chronic sinusitis and viral infections experienced symptom improvement following treatment with anti-inflammatory drugs and IVIg therapy.

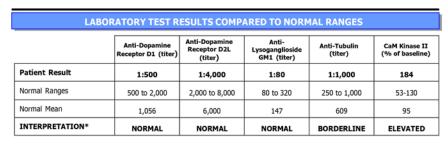


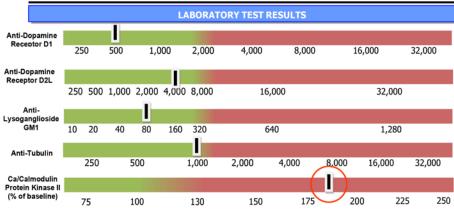
Autoimmune Brain Panel[™] test results (formerly known as the Cunningham Panel[™])

PRE-TREATMENT:

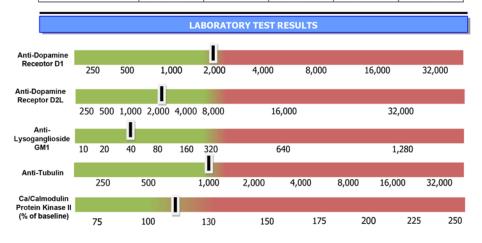
Patient Symptomatic

Elevated Calcium/calmodulin-dependent protein kinase II (CaMKII) 184 consistent with sensory and motor abnormalities, emotional lability, separation anxiety, regression, sleep disturbances, dysgraphia, aggressiveness, and choreiform movements. Borderline anti-Tubulin autoantibodies 1,000 consistent with obsessive-compulsive symptoms and poor concentration.





LABORATORY TEST RESULTS COMPARED TO NORMAL RANGES					
	Anti-Dopamine Receptor D1 (titer)	Anti-Dopamine Receptor D2L (titer)	Anti- Lysoganglioside GM1 (titer)	Anti-Tubulin (titer)	CaM Kinase II (% of baseline)
Patient Result	1:2,000	1:2,000	1:40	1:1,000	113
Normal Ranges	500 to 2,000	2,000 to 8,000	80 to 320	250 to 1,000	53-130
Normal Mean	1,056	6,000	147	609	95
INTERPRETATION*	BORDERLINE	NORMAL	NORMAL	BORDERLINE	NORMAL



POST-TREATMENT:

Symptom Resolution

Normal CaMKII (113) along with borderline Dopamine 1 autoantibodies (2,000) and anti-Tubulin autoantibodies (1,000) consistent with resolution of symptoms.



The Autoimmune Brain Panel™

An Autoimmune Encephalopathy & Basal Ganglia Encephalitis Panel

The Autoimmune Brain Panel™ comprises a series of high complexity blood tests that assists clinicians in identifying an underlying autoimmune etiology for various neurologic (i.e., involuntary movements, seizure-like episodes, tics) and/or psychiatric (i.e., OCD, severe anxiety, mood swings) manifestations.

The Panel measures elevations in autoantibodies directed against specific targets (Dopamine D1 & D2 receptors, Lysoganglioside, Tubulin) in the basal ganglia region of the brain, which may indicate an autoimmune basis for the patient's symptoms. A cell stimulation assay (CaMKinase II) measures the ability of these autoantibodies to stimulate activation of the CaMKII enzyme which can also indicate the likelihood of the presence of an active infection or re-occurring/reactivated infection.

Elevated levels on one or more of these tests indicate that a person's neuropsychiatric symptoms may be due to a treatable autoimmune disorder (potentially triggered by an infection(s), rather than a classic neurologic or psychiatric illness.

Test results can support a clinician's diagnosis with laboratory evidence of an underlying autoimmune etiology and assist clinicians in selecting an appropriate treatment regime. Studies have found that with proper treatment, a patient's symptoms can be greatly reduced or completely resolved.

Elevated levels on one or more of these tests indicate that neurologic and/or psychiatric symptoms may be due to a treatable autoimmune disorder.

Ordering the Autoimmune Brain Panel™

Moleculera Biosciences can accept authorized clinician test orders from all 50 states in the U.S., as well as other countries. (Note: New York patients must have their blood collected outside the State of New York.) *The Panel can only be ordered by licensed clinicians.* Click below to learn how to order the Autoimmune Brain Panel.

Ordering The Panel

