Abstract 6

Do Systemic Inflammation and Blood-Brain Barrier Failure Play a Role in Pediatric Psychosis?

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Context: Blood-brain barrier (BBB) failure occurring downstream of inflammatory processes plays a role in seizure disorders and other neurological diseases. Human and animal studies have suggested an underlying inflammatory mechanism for a variety of neurological disorders, including schizophrenia. To date, all available reports focused on adult patients with chronic schizophrenia. No studies have evaluated a possible link between inflammation, the BBB, and psychotic events in children or adolescents.

Objective: We wished to test the hypothesis that first-episode psychosis, a prodromic event often leading to chronic schizophrenia, is associated with inflammation and BBB leakage. **Patients:** We studied patients admitted to a pediatric inpatient psychiatric unit. Patients (n = 86) had new-onset psychosis diagnosed using DSM-IV TR criteria for Psychosis NOS, schizophreniform disorder, or schizoaffective disorder. Patients were matched for age, race, and gender with nonpsychotic inpatient controls within the same unit (n = 86). We also compared these values to normal control ranges. An additional 10 psychotic patients and as many normal controls were used for cytokine and S100 β serum level analysis.

Main Outcome Measures: In this study, we measured cellular and serum markers of systemic inflammation and BBB leakage.

Results: White blood cell values revealed a significant increase in absolute monocytes (0.62 \pm 0.29; *P* < .01) and lymphocytes (2.51 \pm 0.8; *P* < .05) in psychotic patients compared to nonpsychotic controls (0.47 \pm 0.16 and 2.21 \pm 0.69, respectively). All other hematologic values were similar between the groups. In addition, psychosis was characterized by increased serum levels of S100 β , a peripheral marker of BBB damage. Several inflammatory mediators (eg, TNF- α , IL1- β , IL-6) were elevated in psychotic children.

Conclusions: These results strongly support a link between systemic inflammation, subsequent BBB failure, and first-episode psychosis in pediatric patients.