An Unusual Cause of Neonatal Seizures in a Newborn Infant

ABSTRACT. Neonatal seizures in the neonatal period are symptoms of numerous underlying disorders of the neonate. We present a case in which neonatal seizures due to cerebral infarction led to a diagnosis in the mother. Neonatal convulsions caused by cerebral artery thrombosis is relatively rare in the neonatal period and is often secondary to indwelling intravascular catheters that cause thromboembolism, but may be associated with many conditions. Cerebral artery thrombosis in newborns, in which antiphospholipid antibodies (APA) were found in the mother, has been described in three case reports. Two of these premature infants were born with other risk factors for thrombosis. APA could not be identified in any of these three infants. In the two cases reported by Silver et al the diagnosis was made several months after birth. This case is unique in the fact that no other risk factors for thrombosis could be identified to explain the infarction, and that APA were found in the offspring of an apparently healthy mother. Whether the prior fetal death was caused by APA remains unclear. The finding of lupus anticoagulant in her child led to the diagnosis of antiphospholipid antibody syndrome in her. We believe that in case of cerebral artery thrombosis in a neonate, with no trivial cause such as an indwelling catheter or sepsis, both mother and infant should be tested for presence of APA, even when the mother seems healthy."

CASE REPORT

A 3-day-old male infant was referred to the neonatology ward because of convulsions. He was born at 39-weeks gestation, after a normal vertex delivery. His 25-year-old mother had a history of one intrauterine fetal death at 20 weeks. This pregnancy had been uncomplicated, although her mother had noticed numerous periods of subtle movements in the last 4 weeks of her pregnancy that felt different from normal fetal movements or uterine contractions and never lasted more than 2 minutes. The infant weighed 3500 g (P50), was 54 cm long (P90), and had a head circumference of 35 cm (P50). A few hours after birth, he started to have multiple subtle seizures manifested by repetitive blinking, smacking movements, right-sided twitching of the head and ipsilateral hand and foot. None of the seizures lasted more than 3 minutes. Neurologic examination was otherwise normal. Hematological evaluation, glucose, and electrolytes were normal. There were no signs of an intrauterine infection. An electroencephalogram showed an intermittent irritative disturbance in the left hemisphere.

An ischemic left-middle cerebral artery infarct with a minor hemorrhage could be identified on magnetic resonance imaging (MRI) (Fig 1). On additional magnetic resonance angiography no vascular aberrations of the large vessels were seen. Duplex of the carotid arteries and transcranial Doppler did not show any abnormalities. Coagulation studies revealed lupus anticoagulant (LAC) immunoglobulin G (IgG) presence, the titer being 18.9 GPL units/mL (normal ≤10 GPL units/mL). Anticardiolipin antibody was negative. Detection of LAC and anticardiolipin IgG was performed according to the methods described by Brandt et al. Activated partial thromboplastin time and prothrombin time were both normal. Protein S, protein C activity, and antithrombin III activity were normal for age. Factor V Leiden was absent. An electrocardiogram showed no signs of heart blockage. His mother was additionally tested (6 days later) for antiphospholipid antibodies (APA) and she seemed to have an anticardiolipin IgG titer of 20 GPL units/mL (normal ≤10 GPL units/mL); LAC was negative. The child was given phenobarbital on day 4. The convulsions were not seen after day 5, and no other problems were encountered. He was discharged on day 8, and is still doing well. After 1 month, he became LAC-negative. At the age of 3 months, the MRI showed an old infarct of the left middle cerebral artery with some cortical atrophy (Fig 2). At neurologic examination at the age of 1 year, no asymmetry in motor function was found and there were no signs of developmental delay.

COMMENTS

In the neonatal period, convulsions are usually attributable to perinatal asphyxia, metabolic derangements, or infections. Thromboembolism is a relatively rare cause of convulsions in the newborn and is often secondary to indwelling intravascular catheters, but may be associated with many conditions. Thromboembolism in newborns of LAC-positive mothers has been described. Affected women characteristically have poor pregnancy outcomes that may be improved with prednisone and low-dose aspirin treatment. Zurgil showed transplacental transfer of APA in 18 pregnant women, but in none of the cases could clinical manifestations of APA syndrome be detected. Titers of APA in affected women may fall after pregnancy. This may explain why anticardiolipin antibodies, and not LAC, could be detected in the mother at the time of determination.

Cerebral artery thrombosis in newborns, in which APA were found in the mother, has been described in three case reports. Two of these premature infants were born with other risk factors for thrombosis. In none of the three cases could APA be identified in the infant. In the two cases reported by Silver et al the diagnosis was made several months after birth. This case is unique in the fact that no other risk factors were identified to explain the brain lesion.

ABBREVIATIONS. MRI, magnetic resonance imaging; LAC, lupus anticoagulant; IgG, immunoglobulin G; APA, antiphospholipid antibodies.
factors for thrombosis could be identified to explain the infarction and that APA were found in the offspring of an apparently healthy mother. Whether the prior fetal death was caused by APA remains unclear. The finding of LAC in her child led to the diagnosis of antiphospholipid antibody syndrome in her. This is comparable with diagnosing systemic lupus erythematosus in mothers of children with Ro-SSA positive congenital heart block or neonatal thrombocytopenia.¹⁰

Fig 1. MRI image of the brain made 1 week postpartum showing edema of the left hemisphere with a small hemorrhage, corresponding with a left middle cerebral artery infarction.

Fig 2. MRI image made at 3 months after birth showing focal cortical atrophy of the left parietal lobe, suggesting an old infarction of the left middle cerebral hemisphere.
We believe that in the case of thromboembolism in a neonate, with no trivial cause such as an indwelling catheter or sepsis, both mother and infant should be tested for the presence of APA, even when the mother seems healthy.

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