COMMENT. Molecular genetic testing is important in diagnosis and therapy of infants with CMS. The administration of esterase inhibitors in patients with COLQ mutations can result in serious complications, and an initial short-term beneficial effect may be misleading. Ephedrine (2 to 3 mg/kg/day) was the most effective therapy in the above study. Most patients with COLQ mutations are disabled from infancy, and muscle weakness is progressive and complicated by ventilatory insufficiency and scoliosis. Clinical diagnosis is supported by repetitive CMAP, and increased muscle weakness following administration of pyridostigmine. COLQ gene mutations are the third most common cause of CMS, and occur as frequently as DOK7 mutation cases.

HISTOCHEMICAL ABNORMALITIES IN VARIOUS FORMS OF CONGENITAL MUSCULAR DYSTROPHY

A large Australasian cohort of patients with congenital muscular dystrophy (CMD) was screened to determine the frequency of various forms, in a study at Children’s Hospital at Westmead; the University of Sydney; University of Melbourne, Australia; and University of Nevada, Reno; and University of Illinois, Chicago. Of 101 patients, 45% were screened by immunofluorescence and showed abnormal staining for glycosylated-a-dystroglycan (DG) in 25%. Half of these had reduced DG by Western Blot test. All patients with abnormal DG staining had DNA sequencing of the fukutin-related protein, fukutin, POMGnT1 and POMT1 genes, and mutations were identified in one patient for each of the genes. Abnormalities in collagen VI immunofluorescence were identified in 12% of CMD patients, COL6 mutations in 8 of 9 patients tested, and laminin a2 deficiency occurred in 8% of cases. (Peat RA, Smith JM, Compton AG et al. The diagnosis and etiology of congenital muscular dystrophy. Neurology Dec 26, 2007 (Epub ahead of print)).

COMMENT. Various histochemical and DNA sequencing abnormalities are identified in a large cohort of CMD cases. Patients with abnormal glycosylated a-dystroglycan immunofluorescence were most common in this cohort. Other studies have suggested that the cause of 50% of all CMDs is a primary deficiency in laminin a2, and Ullrich CMD is a second most common form. Molecular diagnostic testing is important for genetic counseling and an emerging new era of gene therapy (Rando TA. Get personal with gene therapy for muscular dystrophy. Lancet Neurology 2008;7:196-8).

MRI IN CONGENITAL FACIAL PALSY

Magnetic resonance (MR) findings in a 12-month-old boy with congenital unilateral facial palsy and a 9-month-old girl with atypical Moebius syndrome are reported from the National Center of Neurology and Psychiatry, Kodaira, Japan. In the boy with unilateral palsy, MRI showed an asymmetry of internal auditory channels with absence of the right facial nerve. MRI on the girl with Moebius syndrome showed a slightly hypoplastic
brainstem and lack of visualization of the proximal part of the facial nerves. 3D-constructive interference in steady state (3D-CISS) MRI sequences, with reconstructions perpendicular to the bilateral internal auditory channel, were required to demonstrate facial nerve anomalies. (Sasaki M, Imamura Y, Sato N. Magnetic resonance imaging in congenital facial palsy. Brain Dev Feb 2008;30:206-210). (Respond: Dr M Sasaki, E-mail: massaki@nc np.go.jp).

COMMENT. Three-dimensional constructive interference in steady state MRI sequence is useful in the differential diagnosis of congenital facial palsy. 3D-CISS MRI provides T2-weighted images with high spatial resolution.

MOVEMENT DISORDERS

GENETICS OF EARLY ONSET RESTLESS LEGS SYNDROME

Linkage analysis was performed in a four-generational German family with restless legs syndrome (RLS) affecting 15 of 37 family members, in a study at the University of Lubeck, Germany. Age at onset was in early childhood or adolescence in 9 (60%) cases. Clinical findings included a desire to move the legs, paresthesias, motor restlessness at night resulting in sleep disturbance and daytime fatigue. Several family members had severe psychiatric problems, including depression and personality disorder. The inheritance pattern was autosomal dominant. A new RLS gene locus (RLS3) was identified on chromosome 9 in all of 12 patients tested, and 11 of these carried an additional closely linked RLS locus. (Lohmann-Hedrich K, Neumann A, Kleensang A, et al. Evidence for linkage of restless legs syndrome to chromosome 9p. Are there two distinct loci? Neurology February 2008;70:686-694). (Reprints: Dr Christine Klein, Department of Neurology, University of Lubeck, 23538 Lubeck, Germany. E-mail: christine.klein@neuro.uni-luebeck.de).

COMMENT. A linkage to a new locus (RLS3) on chromosome 9p has been identified in a family with RLS of early onset. Five gene loci have previously been mapped in cases of primary RLS to chromosomes 12q, 14q, 9p, 2q, and 20p. To date, no gene mutation has been found. RLS is primary or secondary. The primary form is highly familial; secondary RLS is often associated with iron deficiency, renal disease, or pregnancy. The pathophysiology may be related to dopamine insufficiency and low iron storage in substantia nigra.

NEUROCUTANEOUS SYNDROMES

LINEAR NEVUS SEBACEUM SYNDROME AND INFANTILE SPASMS

Two infants with linear nevus sebaceum syndrome and infantile spasms are reported from Safra Childrens Hospital, Sheba Medical Center, Tel Hashomer, Israel; and Hospital for Sick Children, Toronto, Canada. Case 1 presented at age 4 months with focal motor and generalized convulsive seizures with low-grade fever. Family history was positive for febrile seizures in the mother. A 3-cm gray-yellow scaly patch was noted on the frontal-central scalp area that enlarged and turned red and thickened after discharge. Brain MRI showed bilateral