P-060 - SANFILIPPO DISEASE IN A BOLIVIAN PATIENT, CARRIER OF UNCERTAIN SIGNIFICANCE VARIANTS FOR OTHER GENETIC PATHOLOGIES

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INTRODUCTION: The Sanfilippo´s disease or mucopolisacaridosis (MPS) 3A occurs in 1/70,000 newborns (NB), is a rare disease with an autosomal recessive pattern besides the enzyme replacement therapy is still in study. The Cornelia de Lange syndrome occurs in 1/45,000 NB and is autosomal dominant. The Ritscher Schinzel syndrome has few reported cases also with an autosomal dominant pattern. These three conditions share in common some facial dysmorphism and generalized developmental delay. OBJECTIVE: To describe the case of a bolivian patient with Sanfilippo disease, carrier of others genetic variants. MATERIAL AND METHODS: Male, 4 years old, product of 1st pregnancy of non-consanguineous parents, mother and father of 21 years. Without exposure to teratogens, he was born full-term, eutocic delivery, without significant family history. Pathological antecedents of psychomotor development delay and behavior disorders. Physical examination show normal head circumference, weight and height, coarse facies, facial hirsutism, low-set auricular implantation, horizontal palpebral fissures, thick lips. Neck and thorax short, dorsal kyphosis. Ataxic March, language absent. No cardiac or ophthalmological pathology. RESULTS: Positive Berry test, Enzymatic dose arylsulfatase B, alpha iduronidase, Idurnate sulfatase normal. Exoma: Gen SGSH, exon 6 c.703G>A p.(Asp235Asn) corresponding to missense mutation, pathogenic variant, in silico deleterious, and Intron 6 c.745+1G>C corresponding to substitution, probably pathogenic variant, in silico without previous report. Gene SMC3, exon 11 c.874A>T p.(Ser292Cys) corresponding to missense mutation, variant of uncertain significance (VUS), tolerated in silico. Gen CCDC22, exon 4 c.421C>A p.(Pro141Thr) corresponding to missense mutation, variant of uncertain significance, in silico deleterious. DISCUSSION: This is a compound heterozygote for Sanfilippo disease, with a pathogenic variant and another probably pathogenic. The coexistence of two other VUS in relation to the Cornelia de Lange syndrome and Ritscher Schinzel whose facial features are striking. This case could exemplify the epistatic action of the reported genes or the effects of modifier genes.