Case Report

Split-Hand Malformation in a 4-Year-Old Child

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1. Introduction

Split-hand malformation (SHM) is a congenital defect resulting from a chromosomal abnormality in the 7q21q22 region leading to abnormal limb morphogenesis [1]. The presentation of SHM could vary from absence of a single finger to the presence of only one finger (monodactyly) [2]. SHM can occur in isolation or as a part of various complex syndromes [3]. Isolated SHM is usually inherited by the autosomal dominant mode [4]. We report a rare sporadic case of isolated SHM most likely caused by de novo mutation and discuss its etiology, pathogenesis, antenatal diagnosis, genetic counseling, and management.

2. Case Presentation

A 4-year-old child presented to the pediatric department with a deformed left hand since birth. The parents of the child had a nonconsanguineous marriage. Our patient was second in birth order. There was no history of similar malformations in the parents, sibling, or close relatives. General and systemic examination of the child appeared unremarkable. On local examination, a median cleft was present on the affected hand.

The left hand showed aplasia of the 3rd and 4th digits, giving a characteristic lobster-claw appearance (Figure 1). The X-ray showed normal metacarpals but absence of phalanges of the 3rd and 4th fingers. Language development was appropriate for his age. Audiometric tests ruled out sensorineural hearing loss. A diagnosis of SHM was reached clinically. The routine blood and urine investigations did not show any abnormalities. Karyotyping revealed normal male 46, XY karyotype. Further molecular genetic studies could not be carried out due to the economic constraints of the patient’s family. The child was discharged following genetic counseling to the parents.

3. Discussion

SHM is one of the milder manifestations of a genetic disorder called split-hand/split-foot malformation (SHFM) [3]. SHFM, depending on varying levels of its manifestations, causes aplasia/hypoplasia of some of the fingers/toes, syndactyly, and presence of median clefts in the affected hands/feet [1–3]. These limb deformities may occur in isolation, as in our case, or in combination with several complex syndromes. SHM, if syndromic, occurs in conjunction with

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ectodermal dysplasia, cleft lip/palate, mental retardation, and sensorineural deafness [2–4].

SHFM accounts for 8–17% of all limb malformations. The incidence of isolated SHFM is estimated to be 1 per 8000 births, of which 80% have only one affected limb with upper limb predominance [5]. Isolated SHM is caused by mutations at 7q21.3-q22.1, dysregulating the DLX5 and DLX6 genes [3, 4]. DLX6 and DLX5 code transcription factors are largely restricted to the apical ectodermal ridge (AER), a specialized region in the ectoderm required for normal limb skeletal development and morphogenesis [6]. An abnormality in the functioning of the AER leads to abnormalities in the differentiation of the central rays which form the 2nd, 3rd, and 4th digits leading to limb anomalies [5, 6].

SHM is inherited predominantly by the autosomal dominant mode with reduced penetrance, although X-linked and autosomal recessive forms have also been reported [3]. Our case is most likely sporadic in nature as none of the parents, siblings, or close relatives of the patient had similar limb malformations. Also, isolated SHM involving only one upper limb is usually associated with sporadic cases. However, molecular genetic studies as Fluorescence In Situ Hybridization, array-Comparative Genomic Hybridization, and next generation mate-pair sequencing are essential to characterize and truncate the loci of the chromosomal abnormalities in the patient for publication.

Consent
Written informed consent was obtained from the parents of the patient for publication.

Conflicts of Interest
The authors declare that there are no conflicts of interest regarding the publication of this article.

References