

Association of generalized aggressive periodontitis and ectrodactyly-ectodermal dysplasia-cleft syndrome

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Ectrodactyly-ectodermal dysplasia-cleft (EEC) syndrome is an autosomal dominant disorder characterized by the triad of ectrodactyly, ectodermal dysplasia, and facial clefting. Even though literature has documented the association of various genetic disorders with aggressive periodontitis, the periodontal manifestations in patients with EEC syndrome have never been addressed. This case report presents the periodontal status of three patients in a family with EEC syndrome. The presence of generalized aggressive periodontitis was noticed in these patients. EEC syndrome could be a new addition to the group of genetic disorders associated with aggressive periodontitis.

Key words: Aggressive periodontitis, case report, ectrodactyly-ectodermal dysplasia-cleft syndrome

Introduction

Ectrodactyly-ectodermal dysplasia-cleft syndrome (EEC syndrome) is an autosomal dominant disorder characterized by the triad of ectrodactyly, ectodermal dysplasia, and facial clefting.^[1] Although each defect that comprises the syndrome has been known to occur as a separate entity, the constellation of all the three anomalies appears to be a rare occurrence.^[2] Celli *et al.*^[1] reported that the mutation in p63 gene—a homolog of p53 gene—is found to be associated with EEC syndrome.

p63 α , the predominant isoform in epithelial basal cell layers, is responsible for major anomalies found in patients with EEC syndrome.^[3]

Several case reports^[4-6] of EEC syndrome has been published pertaining to the dermatological, urogenital, and dental/oral manifestations of EEC syndrome. Oral manifestations of patients with EEC syndrome reported so far include cleft lip or palate, cleft palate alone, hypodontia, microdontia, anodontia, xerostomia contributing to high caries rate with dry granulomatous lesions on lips, and parotid duct atresia. The periodontal manifestations in EEC syndrome has not been documented in literature.

Although, literature^[7,8] has documented the association of various genetic disorders with aggressive periodontitis, the periodontal manifestations in patients with EEC syndrome have never been addressed. This case report may be the first in literature to document the periodontal manifestations of three patients in a family with EEC syndrome.

Methods

A 23-year-old female presented to the Outpatient Division, Department of Periodontics, Govt. Dental College, Kozhikode, Kerala, India [Figure 1] with congenital disorders on both limbs [Figures 2 and 3]. She was born as the third of six children of non-consanguineous parents after uncomplicated 37 weeks of gestation. Her 63-year old-father had congenital disorder of hands and feet [Figures 4 and 5]. Her 17-year-old younger brother also had congenital disorder of hands and feet

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10.4103/0971-6866.100793

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[Figures 6 and 7], but her mother and other three siblings were normal. It was observed that the female subject , her

father and younger brother had a genetic disorder termed as ectrodactyly ectodermal dysplasia- clefting syndrome



Figure 1: Female patient with coarse and sparse hair, sparse eyebrow, and hypertelorism

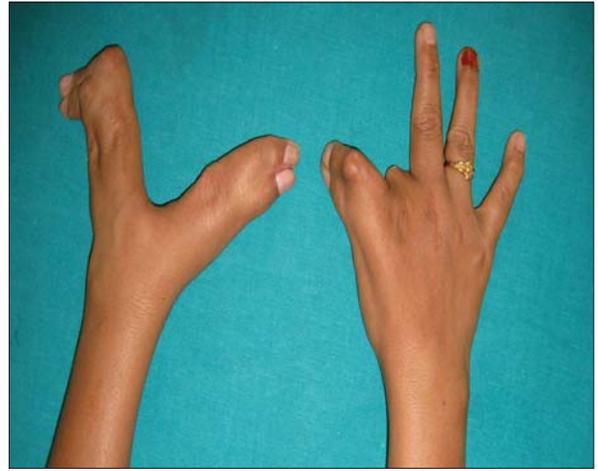


Figure 2: Hands with syndactyly, clinodactyly, and ectrodactyly of female patient



Figure 3: Feet with syndactyly, clinodactyly, and ectrodactyly of female patient

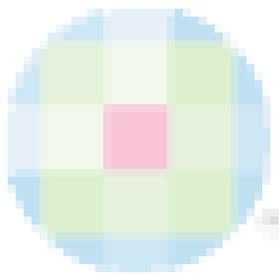


Figure 4: Congenital disorder of hands of the subject's father



Figure 5: Congenital disorder of feet of the subject's father



Figure 6: Congenital disorder of hands of the subject's brother

(EEC) syndrome. She was 155cm tall and weighed 47 kg and had nasal twang in her voice, coarse and sparse hair, sparse eyebrow, hypertelorism, syndactyly, ectrodactyly on both limbs, deep cleft on both limbs, and dysplastic nail of the left toe. Biochemical and hematological parameters, hormonal levels, blood hemoglobin level, and renal function were normal. Intelligence and other developmental milestones were also within the normal limits.

Oral examination

The patient had only 20 teeth [Figure 8] of which mandibular incisors had exfoliated a few months ago, and several other teeth were extracted due to mobility. She had soft palate cleft with missing uvula [Figure 9].

In both the jaws, teeth exhibited pathologic migration and mobility. Probing pocket depth measurements were performed on each tooth using a graduated Williams periodontal probe. Extreme probing depth measurements were noticed at maxillary right and left first molars and mandibular left first molar.

Panoramic radiograph [Figure 10] revealed generalized angular pattern of bone loss with greater loss around maxillary first molars, maxillary central incisors, maxillary second premolar, mandibular left second premolar, and first molar. Based on clinical and radiographic details, a diagnosis of generalized aggressive periodontitis was made.

The subject's father and brother were also diagnosed with generalized aggressive periodontitis.



Figure 7: Congenital disorder of feet of the subject's brother



Figure 8: Intraoral photograph of female patient demonstrating missing teeth



Figure 9: Intraoral photograph of the patient demonstrating soft palate cleft with missing uvula of patient



Figure 10: OPG of the female patient

Discussion

The ectodermal dysplasias are congenital heterogeneous group of inherited disorders with the primary defect in the development of embryonic ectodermal structures. Pure ectodermal dysplasias are manifested by defect in ectodermal structures alone, while ectodermal dysplasia “syndromes” like EEC syndrome have combination of ectodermal defects in association with other anomalies. Neuroectodermal and mesoectodermal derivatives and part of maxillofacial skeleton are also involved in EEC syndrome. This 23-year-old female patient manifested all the three classical features of EEC syndrome, which is a rare occurrence, while her family members (father and brother) showed variable expression of the EEC syndrome without the intraoral clefting. The etiology of EEC syndrome is genetic, and can be both familial and sporadic. Mutation in p63 gene^[1] is found to be associated with EEC syndrome.

Several case reports of EEC syndrome have been published,^[4-6] most of them pertaining to the dermatological, urogenital, and dental manifestations of EEC syndrome. The periodontal manifestations in EEC syndrome patients have never been focused in literature so far. The highly progressive and destructive nature of the bone loss around the teeth (aggressive periodontitis) might have developed at an early in life in these patients. Factors other than pathogenic microorganisms surrounding the tooth in the dental plaque could have influenced the actual clinical presentation. A genetic component might have also cumulated the pattern of periodontal disease in these patients. Quinn *et al.* in 1998^[9] reported that aggressive periodontitis may follow a dominant mode of transmission with the locus for aggressive periodontitis somewhere between the chromosomes number 3 to 7. On the other hand, autosomal recessive mode of transmission has also been noted for aggressive periodontitis.^[10] Taken together, these studies indicate that both etiologic as well as genetic heterogeneity exist for aggressive periodontitis.

The gene for EEC syndrome is also found somewhere between chromosome 3 to 7.^[3] There could be an overlapping association between locus on chromosomal group 3 to 7 for aggressive periodontitis and EEC syndrome, which could have accounted for the aggressive

pattern of periodontal manifestations in these patients. This could open further avenues in investigating the overlapping association of genetics in EEC syndrome and aggressive periodontitis.

The 1999 classification of periodontal diseases and conditions have proposed various genetic disorders associated with periodontitis. Since no other reports exist in literature regarding the periodontal status of patients with EEC syndrome, EEC syndrome could be a new addition to the list of genetic disorders that predispose to periodontal tissue breakdown. It is essential that further research is needed to determine the genetic status of such patients. Management of cases with EEC syndrome and generalized aggressive periodontitis requires a multi-disciplinary approach.

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Cite this article as: Joseph R, Nath SG. Association of generalized aggressive periodontitis and ectrodactyly-ectodermal dysplasia-cleft syndrome. *Indian J Hum Genet* 2012;18:259-62.

Source of Support: Nil, **Conflict of Interest:** None declared.