


CASE REPORT

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Oculo-facio-cardio-dental (OFCD) syndrome: a case report

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Abstract

Background Oculo-facio-cardio-dental (OFCD) syndrome is a rare condition that affects the eyes, face, heart, and teeth of patients. One notable dental characteristic of OFCD is radiculomegaly, or root gigantism, which highlights the role of dentists in detecting this syndrome. OFCD is an X-linked dominant syndrome that results from a variant in the *BCOR* gene. Our study presents the first documented case of OFCD in Vietnam and reports a novel *BCOR* gene variant observed in this case.

Case presentation A 19-year-old Vietnamese female patient with an extremely long root with an abscess was clinically examined for the expression of OFCDs. The radiograph and the variant in *BCOR* gene were also evaluated. We identified abnormalities in the teeth, as well as ocular, facial, and cardiac features, with radiculomegaly of the canines being a specific symptom for OFCDs. The patient's genetic analysis revealed a pathogenic heterozygous deletion at intron 11 of the *BCOR* gene, representing a novel variant.

Conclusion Oculo-facio-cardio-dental syndrome (OFCD) is an extremely rare condition characterized by abnormalities in the eyes, face, heart, and teeth, often caused by variants in the *BCOR* gene. Radiculomegaly, or enlarged dental roots, is a key diagnostic feature of OFCD, and early detection is crucial for preventing future dental complications.

Keywords OFCD syndrome, Dental anomalies, *BCOR* gene

Introduction

Oculo-facio-cardio-dental (OFCD) is a sporadic syndrome (OFCD, MCOPS2; OMIM #300166) involving abnormalities in patients' eyes, face, heart, and teeth. As it was related to dental abnormalities, this condition may often be discovered by dentists.

In 1980, Hayward [1] was the first person to the relationship between a patient's abnormally long teeth and congenital cataracts. In 1990, Marashi and Gorlin [2] reported three similar cases and hypothesized that abnormal root size and congenital cataracts could be characteristics of a specific syndrome. In 1993, Wilkie *et al.* [3] demonstrated a case of a mother and her daughter presenting with ocular, facial, cardiac, and dental abnormalities. He believed that it was an X-linked dominant disorder. Gorlin *et al.* [4] in 1996 named it OFCD syndrome. To date, there were no more than 100 OFCDs

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cases have been reported all over the world. In those reports, the most of patients were female. The study conducted by David Ng has demonstrated that OFCD syndrome is associated with a genetic variant located on the X chromosome [5].

Despite the presence of characteristic symptoms affecting the facial structures and teeth of patients, the rarity of the condition may increase the likelihood of misdiagnosis by physicians as other disorders [6, 7].

As a result, this article aims to summarize symptoms of this extremely rare syndrome and report an OFCD—diagnosed female patient with a discovery of a novel gene variant of *BCOR*. This was the first reported case of this syndrome in Viet Nam.

Case report

The case presents a 19-year-old Vietnamese female patient who came to the Maxillofacial Surgery Department of the National Hospital of Odonto-Stomatology in Ho Chi Minh City with a periapical infection and a fistula that had developed from her left mandibular canine. The patient's past medical history was obtained by interviewing both the patient and her mother, and all patient-specific information was de-identified. Written informed consent was obtained from the patient's legal guardian for the publication of photographs and any accompanying information for the purpose of this case report.

Past medical history

Strabismus was detected at six months old. At the age of four, she was diagnosed with congenital cataracts. She underwent surgery to remove her cloudy lenses and had them replaced with new, artificial lenses (IOLs—Intraocular Lenses). At 14 years old, she was diagnosed with congenital cardiac diseases, including ostium secundum atrial septal defect, pulmonary artery hypertension and 4/4 leaky tricuspid valves. Furthermore, upon examining the patient's mother, we observed the presence of strabismus, even though no evidence of radiculomegaly was found on her panoramic radiograph.

There was no family history of any genetic diseases. The patient has two younger sisters with typical development. The patient's mother reported a fever and was hospitalized for one week at 22 weeks gestation. No abnormalities were detected during her mother's antenatal ultrasounds.

Extra-oral examination

Clinically, the patient presented with a long and broad face, as well as a concave facial profile in the lateral view. Other physical features included thick eyebrows,



Fig. 1 Strabismus, microphthalmia on her left side



Fig. 2 Bifid nasal tip

a broad and protrusive mandible, and eye abnormalities such as strabismus and microphthalmia on the left side. Visual impairment was present in both eyes, and there were no signs of secondary glaucoma. Additionally, a broad nasal tip with separation of anterior cartilage nasal was observed, along with a fistula on the left chin surrounded by an inflammatory area (Figs. 1, 2). There were no signs of hearing impairment, protruding or hypoplastic ears. The patient had normal pronunciation, but presented with intellectual disability.

Limb examination

The patient presented with IV-toe camptodactyly and a I-hammer toe. There were no abnormalities detected in her hands (Figs. 3, 4). She did not have hearing



Fig. 3 Abnormal foot development (IV—toe camptodactyly and a I—hammer toe)



Fig. 4 Long and slender fingers

impairment, protruding ears, or hypoplastic ears. Her pronunciation was normal, but she had intellectual disability. Radiograph examination revealed IV-toe camptodactyly and I-hammer toe, and her hand radiograph did not show any abnormality (Fig. 5).

Oral examination

The patient had a narrow, high-arched palate, with severely crowded teeth and multiple dental caries. Specifically, teeth 11, 12, 16, 21, 26, 32, 33, 36, 43, 46, and 47 had significant decay and required further dental treatments, including extractions and endodontic procedures. Additionally, glossitis was present (Fig. 6).

Radiograph findings

The Panoramic radiograph revealed a condition of severe crowding of the teeth with malposition of teeth 34, 44, retained roots of teeth 11, 12, 21, 26, 32, 33, 36, 43, 46 with many dental decays. The impacted tooth 24 was also noted. Additionally, the lower premolars and upper right second premolar showed signs of radiculomegaly. Furthermore, the lower anterior teeth had very long roots, and the 37 appeared taurodontic. The 48 was

under-developed for 19 years of age. Notably, elongated canines' roots with open apices were demonstrated in all quadrants of her jaws. Moreover, a periapical infection was detected on her left mandibular canine through the panoramic radiograph (Fig. 7).

Genetic analyses

Genomic DNA was extracted from peripheral blood using the GeneJET™ Whole Blood Genomic DNA Purification Mini Kit (Thermo Fisher Scientific, MA, USA). Genomic DNA was used to amplify exons 1 through 15 and the exon–intron boundaries of the *BCOR* gene using pairs of PCR primers designed by our own. Purified PCR products were sequenced in both directions using Big Dye Version 3.1 and an ABI 3500 Genetic Analyzer (Applied Biosystems, CA, USA). As a result, the patient was found to be heterozygous for a novel single-base deletion within intron 11 (IVS11-2delA) of the *BCOR* (Fig. 8).

Diagnosis and treatment

The patient was diagnosed with periapical infection resulting from her left mandibular canine, and was found to have OFCD syndrome. The treatment plan included a root canal treatment for the affected tooth followed by an extraoral apicoectomy to remove the fistula on her left chin (Figs. 9, 10).

Follow up

The patient experienced no pain or discomfort in the surgical area. An OPG was taken two weeks post-surgery, and no abnormalities were observed on the X-ray. The patient was re-examined 4 months after surgery and reported no chin pain. At that moment, we did not conduct an X-ray. No adverse events were documented.



Fig. 5 Foot X-ray and hand X-ray



Fig. 6 Intra examination showed high arched palate, many dental decays, and mal-occlusion with malposition of teeth 33, 34, 44, 25, heavy dental plaque, stomatitis, glossitis, and bifid uvula was also seen



Fig. 7 Panoramic panoramic radiograph revealed seriously crowding teeth, many dental caries and radiculomegaly canine with open apices

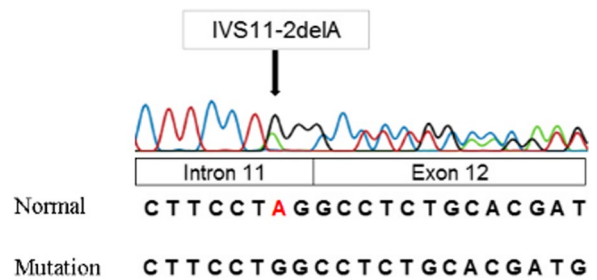


Fig. 8 *BCOR* gene sequencing results: detected heterozygous variant IVS11-2delA on intron 11 of *BCOR* gene



Fig. 9 Panoramic film after root canal treatment

Discussion

OFCD is a rare genetic syndrome with no more than 100 cases reported in literature [8–10], characterized by ocular, facial, cardiac, and dental abnormalities. Early and definitive diagnosis of this syndrome is considered problematic. However, OFCD syndrome has some specific dental symptoms, including an extreme elongation of canine roots and an open apex. Fortunately, these signs could be easily identified on panoramic dental radiograph.

OFCD syndrome has been misdiagnosed as maternal exposure to rubella during pregnancy [6, 11, 12]. However, in some cases, abnormal symptoms in the teeth, particularly radiculomegaly, and the absence of maternal rubella infection during pregnancy, help to differentiate the diagnosis [13]. In addition to the four main symptoms (including ocular, facial, cardiac, and dental), OFCDs patients often have the abnormality in the ears (hearing impairment, protruding ears, hypoplastic ears) [10, 12, 14], extremities (toe syndactyly, hammer toes, radioulnar synostosis) [6, 10, 15] and mental retardation [6, 10, 16]. Numerous cases with several BCOR variants have been reported in the medical literature [8, 9, 13, 14, 17–27] (see Table 1). In our report, the patient’s chief complaint

was the appearance of a fistula developing from her left mandibular canine. She was then indicated to take a panoramic radiograph, and we found that all her canine teeth had extreme elongation with their apex opening. Clinical examination and medical history demonstrated that this patient had abnormalities in her eyes (congenital cataract, cross-eyed), her face (long and narrow faces, bifid nasal tip) (Fig. 2), and her heart (ostium secundum atrial septal defect, pulmonary artery hypertension, and 4/4 leaky tricuspid valves). We speculated that this patient might have an OFCD syndrome. Therefore, we decided to take a screening for BCOR variants. The result showed that a heterozygous deletion variant IVS 11-2delA was detected in intron 11 of gene BCOR, a novel variant. With these above symptoms, this patient was planned to receive thorough treatment, including extracting all her root teeth, filling all decayed teeth, undergoing a root canal treatment for the left mandibular canine, and having an extraoral apicoectomy with a fistula removal on her left chin. OFCD is a rare syndrome affecting many organs. Early identification of this disease helps prevent oral complications and endocarditis progression due to caries teeth. As a result, we could combine a wide range of specialties to provide patients with comprehensive care.

Conclusion

Radiculomegaly is a crucial dental symptom that is highly specific to OFCDs. This syndrome is often detected by dentists during dental panoramic radiograph examinations. Therefore, dentists play an essential role in identifying and diagnosing this OFCD syndrome. Dentists should identify patients who may have the condition and refer them to geneticists for further examination and testing. In other words, patients need an accurate diagnosis to receive prophylactic treatment for other related conditions.



Fig. 10 Extraoral apicoectomy with a fistula removal

Table 1 Literature review of case report with BCOR variant

| General information | Current study | | | | | | | | | |
|-----------------------|-----------------------------------------------------------------------------------------------------------|-----------------------|-----------------------|-------------------------------------|---------------------------------|-----------------------------------|-----------------------------------------------------------------------------------------------------|-------------------------------------|------------------------------------------------|-----------------------------------------|
| | Ragger/2018/USA [9] | Case 1 | Case 2 | Case 3 | Case 4 | Morgan/2019/USA [18] | Zhang/2019/China [19] | Song/2019/Korea [20] | Tsuwaki/2005/Japan [13] | Di-Stefano/2015/Italy [14] |
| Age (year) | 19 | 13 | 21 | 3 | 10 | 10 | 4 | 31 | 16 | 2 |
| Gender | Female | Female | Male | Female | Female | Female | Female | Female | Female | Female |
| BCOR variant | IVS11-2delA (c.4596-2delA) | c.2428C>T p.(Arg810*) | c.254C>T p.(Pro85Leu) | c.1209_1210delCC p.(Gln404Alafs*35) | Exon 4 c.265G>A | c.2514del(G) | c.1296delT | n/a | n/a | a de novo heterozygous delXp11.4 |
| Inheritance | De novo | De novo | Maternal | De novo | Denovo | De novo | De novo | De novo | | |
| Ocular | | | | | | | | | | |
| Microphthalmia | Left side | Bilateral | Bilateral (severe) | Bilateral | | Left eye microphthalmia | | | Bilateral | Right eye |
| Anophthalmia | No | | | | | | | | | |
| Congenital cataract | Bilateral | Bilateral | | Bilateral | Bilateral | + | + | + | Bilateral | |
| Glaucoma | n/a | Unilateral | | | | In the right eye | | | + | + |
| Strabismus | + | | | | | | | | | |
| Posterior embryotoxon | n/a | | | | | | | | | |
| Other | Strabismus, left side | | | | Artificial eye on the left side | Hemangioma near her right eyebrow | Bilateral ptosis, eyebrows curvature; mixed nystagmus; right eye exohypertropia in primary position | | | Secondary cataract |
| Craniofacial | Long, broad face, concave facial profile in a lateral view, broad and protrusive mandible, prominent chin | | | | | | | | | |
| Midface hypoplasia | + | | + | | | | | | | |
| Nasal anomalies | Broad nasal tip with separation of anterior nasal cartilage, bifid nasal tip | | + | | Broadening of the nasal tip | | Broad nasal tip | Nasal tip was prominent and bulbous | Broad nasal tip separated nasal cartilage | Broad nasal tip, depressed nasal bridge |
| Ear anomalies | n/a | | + | + | Preauricular tag | | Protruding ears | No | Hearing on the right side is impaired slightly | Depressed nasal bridge |
| Cleft palate | Bifid uvula | | | - | | | + | no | no | |

Table 1 (continued)

| Current study | | Ragge/2018/USA [9] | Kato/2018/ Japan [17] | Morgan/2019/ USA [18] | Zhang/2019/ China [19] | Song/ 2019/ Korea [20] | Tsuwaki/2005/ Japan [13] | Di-Stefano/2015/ Italy [14] | | |
|-----------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------|-----------------------------|-----------------------------------|-------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------|-----------------------------------------------|---------------------------------------|----------------------------------------------------|--------------------------------------------------------------------|
| General information | Current case | Case 1 | Case 2 | Case 3 | Case 4 | Case 5 | Case 6 | Case 7 | Case 8 | Case 9 |
| High arched palate | + | | | + | | | + | | | |
| Other | Thick eyebrows, narrow palate, and mandible, small mouth | Downslanting palpebral fissures, long face, tall forehead, thick eyebrows, Long Philtrum | | | Elongated, biprotrusive, with a thick lower lip | | Flat and slightly long, dovetail-shaped uvula | | | Broad forehead, bifid uvula |
| Cardiac | | | | | | | | | | |
| ASD | Ostium secundum atrial septal defect | | | | | + | no | + | | + |
| VSD | n/a | | | | | | no | | | + |
| Other | Pulmonary artery hypertension, 4/4 leaky tricuspid valves | | Triple heart sounds | | | | | | | Patent ductus arteriosus (PDA), persistent left superior vena cava |
| Dental | | | | | | | | | | |
| Late eruption of first teeth | n/a | + | | + | n/a | | | | | |
| Impacted teeth | Tooth 24 | | | | Tooth 23 was retracted and extruded | | | | + | |
| Delayed loss of primary dentition | n/a | + | | | | | | | | |
| Radiculomegaly | Canines, premolars, and lower anterior teeth | | | | | | | | | |
| Tautodontism | Tooth 37 | | | | | | | Teeth 13, 22, 23, 33, 34, 43, 44 | | |
| Fused incisors | No | | | | | | | | | |
| XQ | | | | | | | | | | |
| Other | Crowding with malposition of teeth 34,44, many dental caries (1,1,2, 16,21, 26,32,33,36,43,46, 47) and periapical infection of tooth 33 with skin leaking | Double row of teeth | Recurrent dental infections | Abnormal crown canines + incisors | Malocclusion, anterior dental crowding, incisors, and first premolars had Open apex | Long roots of her teeth with one missing tooth and first primary tooth loss at 6–7 years of age | Crossbite | Enamel hypoplasia, crown malformation | Lateral crossbite, Oligodontia, enamel hypoplasia, | |

Table 1 (continued)

| Current study | | Ragge/2018/USA [9] | Kato/2018/ Japan [17] | Morgan/2019/ USA [18] | Zhang/2019/ China [19] | Song/ 2019/ Korea [20] | Tsuwaki/2005/ Japan [13] | Di-Stefano/2015/ Italy [14] | | |
|----------------------------|----------------------------------------------------------|----------------------------------------------------------|--------------------------------------|---------------------------------------------------------|------------------------|------------------------|--------------------------------------------------------------------------------------------|-----------------------------|--------------------------------------------------------------|-------------------------------------------|
| Current case | | Case 1 | Case 2 | Case 3 | Case 4 | Case 5 | Case 6 | Case 7 | Case 8 | Case 9 |
| General information | | | | | | | | | | |
| Skeletal | Glossitis, stomatitis, heavy dental plaque, and calculus | | | | | | | | | |
| Hands | Long and slender finger | Long Finger | 5th Finger clinodactyly, long Finger | Long Finger | n/a | n/a | Short | No | | Clinodactyly of the fifth finger |
| Feet | IV-toes camptodactyly, Hammer toes, long toes | Long toes | | | II-Hammer toes | n/a | Hammer-type big toes; flexion deformity (2–4 toes of right foot and 2–3 toes of left foot) | No | I-long and wide toe, hammer-type flexion of toes 2 through 4 | Syndactyly 2–3, ham-mer toe of the second |
| Other | n/a | | Scoliosis | | | | Asymmetry of hand size | | Forearm on the right side was slightly shorter | |
| Developmental | | | | | | | | | | |
| ID | No intellect defect | | | | | n/a | | | | |
| Motor delay | Mental retardation | | + | | n/a | n/a | No | | | No |
| Speech delay | n/a | | + | | n/a | n/a | | | | No |
| MRI findings | n/a | | N | | n/a | n/a | n/a | | | No |
| Lipomatous lesion | n/a | | | | n/a | n/a | n/a | | | |
| Other | | | | Moderate BA, broad lateral ventricles | | | | | | |
| Other findings | | | | | | | | | | |
| GU anomalies | n/a | Urethral hypoplasia, renal dysplasia, renal failure, VUR | Primary enuresis | Cryptorchidism, vesicoureteric reflux, primary enuresis | n/a | n/a | | | | n/a |
| Other | | Hypotonia | Stage III T-cell lymphoma | | | | Bilateral papiloma of choroid plexus (PCP), supratentorial hydrocephalus | | | |

Table 1 (continued)

| General information | Current study | Mc Govern/2006/Ireland [21] | Atiq/2012/USA [22] | Danda/2014/India [23] | Martinho/2019/Portugal [8] | Türkkahraman/2006/Turkey [24] | Verm/2014/India [25] | Zhou/2018/USA [26] | Zhu/2015/China [27] |
|-----------------------|-----------------------------------------------------------------------------------------------------------|-----------------------------------------|--------------------|-----------------------|----------------------------|-------------------------------|----------------------|-----------------------------------|---------------------|
| | | | | | | | | | |
| Age (year) | 19 | 8 days | 39 | 8 | 26 | 15 | 24 | 5 weeks | 7 months |
| Gender | Female | Female | Female | Female | Female | Female | Female | Female | Male |
| BCOR variant | IVS11-2delA (c4596-2delA) | n/a | | c.3490C>T (p.R1164*) | | | | p.R1163X (c.3487 C(T)) | p. R540Q |
| Inheritance | De novo | Maternal | | Heterozygous | | | | | Missense mutation |
| Microphthalmia | Left side | + | | | | | + | + | |
| Anophthalmia | No | | | | | | | | |
| Congenital cataract | Bilateral | | + | + | + | + | + | | |
| Glaucoma | n/a | | | | | | | | |
| Strabismus | + | | | | | | | | |
| Posterior embryotoxon | n/a | | | | | | | | |
| Other | Strabismus, left side | Deep-set eyes, short palpebral fissures | | | | | | | No |
| Craniofacial | Long, broad face, concave facial profile in a lateral view, broad and protrusive mandible, prominent chin | | | | | | | | |
| Midface hypoplasia | + | | | | | | + | | |
| Nasal anomalies | Broad nasal tip with separation of anterior nasal cartilage, bifid nasal tip | | | | | | | Broad nasal tip flat nasal bridge | |

Table 1 (continued)

| Current study | | Mc Govern/2006/Ireland [21] | Atiq/2012/USA [22] | Danda/2014/India [23] | Martinho/2019/Portugal [8] | Türkkahraman/2006/Turkey [24] | Verm/2014/India [25] | Zhou/2018/USA [26] | Zhu/2015/China [27] | |
|------------------------------|-----------------------------------------------------------|-----------------------------|-----------------------------------------------|-----------------------|---------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------|------------------------------------------------------------|-----------------------------------------------|--------------------------------|
| General information | Current case | Case 10 | Case 11 | Case 12 | Case 13 | Case 14 | Case 15 | Case 16 | Case 17 | Case 18 |
| Ear anomalies | n/a | | | | | | | | Left hearing impairment slightly low-set ears | |
| Cleft palate | Bifid uvula | | | | | | | | | |
| High arched palate | + | | + | | | | | | | |
| Other | Thick eyebrows, narrow palate, and mandible, small mouth | | Extended long canine teeth, and nasal changes | High forehead | High nasal bridge | Class II malocclusion on a Class III skeletal base with a prognathic mandible, increased facial proportions, and facial asymmetry | Long and narrow face, high nasal bridge, broad nasal tip with separated cartilages, and a long philtrum | Deeply set eyes and a broad nasal tip long and narrow face | | |
| Cardiac | | | | | | | | | | |
| ASD | Ostium secundum atrial septal defect | + | - | | | | | | + | |
| VSD | n/a | + | - | | | | | | | |
| Other | Pulmonary artery hypertension, 4/4 leaky tricuspid valves | Pulmonary valve stenosis | Mitral valve prolapse | + | Double outlet right ventricle, pulmonary stenosis | Prolapsed mitral valve | | | + | Patent ductus arteriosus (PDA) |
| Dental | | | | | | | | | | |
| Late eruption of first teeth | n/a | | | + | | | | | | + |
| Impacted teeth | Tooth 24 | | | | | Gummy smile, and crowded teeth | Extremely long roots and open apices | | | |

Table 1 (continued)

| | Current study | Mc Govern/2006/ Ireland [21] | Atiq/2012/ USA [22] | Danda/2014/India [23] | Martinho/2019/ Portugal [8] | Türk Kahraman/2006/ Turkey [24] | Verm/2014/ India [25] | Zhou/2018/ USA [26] | Zhu/2015/ China [27] | |
|-----------------------------------|----------------------------------------------|------------------------------|---------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------|---------------------------------|-----------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------|---------|
| General information | Current case | Case 10 | Case 11 | Case 12 | Case 13 | Case 14 | Case 15 | Case 16 | Case 17 | Case 18 |
| Delayed loss of primary dentition | n/a | | | + | | | + | | | |
| Radiculo-megaly | Canines, premolars, and lower anterior teeth | | | | | + | | | | |
| Tautodontism | Tooth 37 | | | | | | | | | |
| Fused incisors | No | | | | | | | | | |
| XQ | | | | Teethskel-etal class I with severe vertical growth pattern, increased gonial angle, steep mandibular plane with retroclined incisors, and competent lips | | | | Permanent teeth with extremely long roots and open apices. The roots of maxillary canines were in relation with the inferior border of the orbits and the lower canine roots almost reached the lower border of the mandible. The maxillary left central incisor had dilacerated root; all four third molars were congenitally missing | | |

Table 1 (continued)

| Current study | Mc Govern/2006/ Ireland [21] | Case 10 | Atiq/2012/ USA [22] | Case 11 | Danda/2014/India [23] | Case 12 | Case 13 | Martinho/2019/ Portugal [8] | Case 14 | Türkkahraman/2006/ Turkey [24] | Case 15 | Verm/2014/ India [25] | Case 16 | Zhou/2018/ USA [26] | Case 17 | Zhu/2015/ China [27] | Case 18 |
|----------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------|---------|---------------------|---------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------|---------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------|--------------------------------|---------|-----------------------|---------|---------------------|---------|----------------------|---------|
| General information | | | | | | | | | | | | | | | | | |
| Other | Crowding with malposition of teeth 34,44, many dental caries (1,1,12,16,21,26,32,33,36,43,46,47) and periapical infection of tooth 33 with skin leaking | | | | Bifid uvula | | | Numerous missing teeth The first upper left molar, upper right canine, upper left lateral incisor, first upper left premolar, first upper right molar, first lower right molar, and first lower left molar are absent | | | | | | | | | |
| Skeletal | Glossitis, stomatitis, heavy dental plaque, and calculus | | | | | | | | | | | | | | | | |
| Hands | Long and slender fingers | | | | Camptodactyly of the 4th and 5th fingers (right > left), proximally placed thumbs, restricted supination, and pronation of the left forearm, camptodactyly and syndactyly of 2nd and 3rd toes, and sandal gap Elbow radiographs at infancy showed left radioulnar synostosis | | | Misalignment | | | | Fingers are normal | | | | | |

Table 1 (continued)

| | Current study | Mc Govern/2006/Ireland [21] | Atiq/2012/USA [22] | Danda/2014/India [23] | Martinho/2019/Portugal [8] | Türkkahraman/2006/Turkey [24] | Verm/2014/India [25] | Zhou/2018/USA [26] | Zhu/2015/China [27] | |
|---------------------|--------------------------------------------------|----------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------|-----------------------------------------|-------------------------------|----------------------|-------------------------------|--------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------|
| General information | Current case | Case 10 | Case 11 | Case 12 | Case 13 | Case 14 | Case 15 | Case 16 | Case 17 | Case 18 |
| Feet | IV-toes camp-todactyly, I-hammer toes, long toes | | | Sandal gaps, syndactyly and camptodactyly of toes | Sandal gap between the 1st and 2nd toes | Valgus foot | | Syndactly of 2nd and 3rd toes | Right clubfoot, and bilateral 2–3 toe syndactyly | |
| Other | n/a | Misalignment of right second toe | | Short stature was observed (122 cm; < 3 SD) | Short stature (111 cm, < 3 SD) | | | | | |
| Developmental | | No | | | | | | | | |
| ID | No intellect defect | | | | | | + | | + | |
| Motor delay | Mental retardation | | | | | | | | + | |
| Speech delay | n/a | | | | | | | | | |
| MRI findings | n/a | | | | | | | | | |
| Lipomatous lesion | n/a | | | | | | | | | |
| Other | | | | | | | | | | |
| Other findings | | | | | | | | | | |
| GU anomalies | n/a | No | | | | | | | | |
| Other | | Anterior positioning of the anus | Several episodes of hypoglycemia several episodes of mental confusion associated with a blood glucose level of less than 40 mg/dL reactive lymph node rare congenital disorder | | | | Tall stature | Umbilical hernia at birth | | CHD structural brain anomalies Axenfeld–Rieger syndrome, Lenz microphthalmia syndrome, and oculo-facio-cardio-dental (OFCD) syndrome |

BA brain atrophy, ASD atrial septal defect, VSD ventricular septal defect, ID Intellectual delay, Cm centimeter, SD standard deviation, MRI magnetic resonance imaging, VUR vesicoureteric reflux, CHD congenital heart defects, BCOR variant The BCOR gene is responsible for coding the BCL6 corepressor protein, N/a not applicable or not available, PDA patent ductus arteriosus, N normal

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Author contributions

NTT contributed significant contributions to the manuscript's acquisition, interpretation, and writing. TTHA, HAV, HVD, NVT, LTC, DTPD, LHNM contributed significant contributions to data interpretation and paper revision. NTT, DTPD, LHNM contributed significantly to the conception/design, acquisition, and interpretation of data, as well as extensively revising the manuscript. The final version was reviewed and accepted by every writer.

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Availability of data and materials

The authors affirm that all data available to sustain the report's interpretations were included in the publication and its supplementary materials.

Declarations

Ethics approval and consent to participate

All approaches were conducted in accordance with applicable rules and regulations as detailed in the Declaration section.

Consent for publication

Written informed consent was obtained from the patient's legal guardian for publication of this case report and any accompanying images. A copy of the written consent is available for review by the Editor-in-Chief of this journal.

Competing interests

The authors declare that they have no competing interests.

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