

Papillon-Lefèvre Syndrome: A Report of Three Siblings

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Abstract We reported an 18 year old Sudanese female patient, Gaali tribe, and another 2 males, brothers ages 10 and 8 years old, presented with diffuse transgradient palmoplantar keratosis with scaly, non-erythematous, psoriasiform-like transgradient plaque in elbows and knees, and periodontitis, the female patient has been misdiagnosed before as a case of Psoriasis and given phototherapy in the form of PUVA with satisfactory results and the 2 brothers has been also missed as a cases of eczemas with poor response to treatment and frequent relapses. The female case is associated with thyroid swelling and increased Thyroid Stimulating Hormone TSH, and considered as a first cases record at Sudan, as Papillon-Lefèvre syndrome is known as a rare presentation globally and in Sudan specifically.

Keywords Syn. Papillon-Lefevre Syndrome, Disease, Papillon-Lefevre, Papillon Lefevre Disease, Papillon Lefevre Syndrome, Keratosis palmoplantar-periodontopathy

1. Background

The Papillon-Lefevre syndrome is psoriasiform hyperkeratosis of the palms and soles that can overflow onto the dorsal surfaces of the hands and feet. Psoriasiform lesions can also be seen on the limbs. This keratoderma associates, as early as infancy, intense gingivitis with alveolar bone lysis and early loss of baby teeth. During childhood, this phenomenon of periodontal disease recurs with rapid loss of adult teeth. This syndrome is accompanied, in half the patients, by enhanced susceptibility to infections (furunculosis, skin abscesses, hidradenitis suppurativa). Anomalies of chemotaxis and phagocytosis by polymorphonuclear leukocytes have been observed (Itin and Fistarol).

This condition is inherited in an autosomal recessive fashion. The frequency of PPK with periodontitis is 4 cases per 1 million population. The male-to-female ratio is equal. Onset occurs between the first and fifth years of life. A variant, Haim-Munk syndrome features, in addition to PPK and periodontitis, arachnodactyly, acroosteolysis, and

onychogryphosis (Ratnavel and Griffiths, 1997).

Clinically, diffuse transgradient PPK may be observed, typically developing within the first 3 years of life. Punctiform accentuation, particularly along the palmoplantar creases, may be seen. Unless treated, periodontitis results in severe gingivitis and loss of teeth by age 5 years. No significant correlation has been demonstrated between the level of periodontal infection and the severity of skin affections, which supports the concept that these major components of this syndrome are unrelated to each other. Patient's exhibit increased susceptibility to cutaneous and systemic infections. Scaly, psoriasiform lesions are often observed over the knees, elbows, and interphalangeal joints. Finally, patients may have malodorous hyperhidrosis (Griffiths et al., 1998).

- ▶ anodontia/**oligodontia** (Very frequent sign).
- ▶ autosomal recessive inheritance (Very frequent sign)
- ▶ dysplastic/thick/**grooved** fingernails (Very frequent sign)
- ▶ hamartoma/tumours of the mouth (Very frequent sign)
- ▶ **palmoplantar hyperkeratosis** (Very frequent sign)
- ▶ **teeth anomalies** (Very frequent sign)
- ▶ arachnodactyly (Frequent sign)
- ▶ intracranial calcifications (Frequent sign)
- ▶ thin/hypoplastic/**hyperconvex** fingernails (Frequent sign)

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Published online at <http://journal.sapub.org/ajdv>

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- ▶ decreased body hair (Occasional sign)
- ▶ increased body hair (Occasional sign)
- ▶ skin tumours/lumps (Occasional sign)
- ▶ sparse/absent scalp hair(generalized) (Occasional sign)

Histologic findings include hyperkeratosis with irregular parakeratosis and moderate perivascular infiltration. Electron microscopic features include lipid like vacuoles in corneocytes and granulocytes, a reduction in tonofilaments, and irregular keratohyalin granules (Griffiths *et al.*, 1998).

Molecular biology findings include mutations in the Cathepsin C gene, mapping to 11q14-q21, which are responsible for this syndrome. Cathepsin C is a lysosomal protease known to activate enzymes that are vital to the body's defenses (Hart *et al.*, 1998; Kimyai-Asadi *et al.*, 2002).

Treatment includes oral retinoids for the PPK. Elective extraction of involved teeth may prevent excess bone resorption. Appropriate antibiotic therapy may be required for periodontitis and recurrent cutaneous and systemic infections. Treatment with acitretin starting at an early age shows promise towards allowing patients to have normal adult dentition (Saywell C and WAD, 1994).

Retinoids slow the alveolar bone lysis and attenuate the palmoplantar keratoderma. This genetic disease is transmitted by autosomal recessive inheritance. It is linked to an almost total loss of Cathepsin C activity (Hart *et al.*, 1999; Toomes *et al.*, 1999).

2. The Case 1

A female patient, student, 18 years old, single, resident in Al-Dqba, north of Sudan. The patient is complaining of dry painful fissured, non-itchy, soles of feet with difficulty on walking since age of 4 months, loss of teeth with swollen painful gum since the age of 15 years old.

Patient's general condition looks well with average intelligence, a thyroid swelling is noticed, uniformly diffuse, moves with deglutition, no regional lymphadenopathy. The condition started at infancy, age of 4 months with peeling of soles of feet, with gradual onset and progressive course, where thickening of both soles ensues, fissured and worse with age and focal thickening in both palms. Patient noticed inability to move some of her hand fingers since childhood. Patient has been diagnosed as a case of Psoriasis and received topical treatment in the form of PUVA with more or less satisfactory result.

Family history: Parents are first degree relatives, **no** similar condition in family, and **heel fissurings** in her cousins is noticed.

On examination: Patient has normal facial features with diffuse transgradient palmoplantar keratoderma, and distant psoriasiform plaques-like at both knees and elbows. Hyperkeratotic knuckle pads mainly at proximal interphalangeal joints, with joint block at both little fingers,

where patient can't approximate palms of both hands together.

Plaques are well defined, demarcated scaly, no fissures or erythema, 1-2 cms in diameter. No hyperhidrosis is seen. Lesions are **skin colored**.

The lesions are **Firm** on palpation.

Palms & soles: Diffuse transgradient palmoplantar keratoderma. **No** Palmar or plantar pits or erythema, as well as arachnodactyly or pseudoainhum of the digits, **not** detected.

Nails: Nail dystrophy with hyperconvexity, transverse and longitudinal ridging is noticed at most of dysplastic nails.

Hair: No Abnormality Detected, with normal body and scalp hair.

Eye: No abnormality seen.

Oral mucosa & Lips: Periodontosis with loss of teeth-Oligodontia. **No** Leukokeratosis.

3. Investigations Done

1. Skin biopsy:

Macro: 2 small pieces of soft tissue, submitted all in one block.

Micro:

Sections show tumor cells form cords and clusters of cells with **basophilic** ovoid nuclei and little cytoplasm.

The cells at the periphery of the cords show **palisading**.

There are a melanin pigments and mitotic figures.

Diagnosis: Basal Cell Carcinoma, BCC.



Figure 1. Diffuse plantar keratoderma both feet



Figure 2. Focal palmar keratoderma both hands



Figure 3. Blocked joints little fingers of both hands



Figure 4. Longitudinal nail ridging of middle finger with ragged nail cuticle



Figure 5. Psoriasis like lesions both knees



Figure 6. Complete denture

2. X-RAY Report:

To show:

Skull:

No intracranial Calcification seen.

Cystic changes of the body of the mandible.

Normal Sella Tursica, no bridging is evident.

No evidence of Increased Intra Cranial Pressure I.I.C.P.

Thyroid function tests:

Results have shown;

FSH raised 15.3

4. The Case 2

A male patient, student, 10 years old, resident in Al-Duba, north of Sudan. The patient is complaining of dry painful fissured, non-itchy, soles of feet with difficulty on walking since age of 4 months, loss of teeth with swollen painful gum since the age of 15 years old.

Patient's general condition looks well with average intelligence.

The condition started at infancy, age of 4 months with peeling of soles of feet, with gradual onset and progressive course, where thickening of both soles ensues, fissured and worse with age and focal thickening in both palms. Patient noticed inability to move some of her hand fingers since childhood. Patient has been diagnosed as a case of Psoriasis and received topical treatment in the form of PUVA with more or less satisfactory result.

Family history: Parents are first degree relatives, 2 similar condition in family, oldest sister and a younger brother.

On examination: Patient has normal facial features with diffuse transgradient palmoplantar keratoderma, and distant psoriasiform plaques-like at both knees and elbows. Hyperkeratotic knuckle pads mainly at proximal interphalangeal joints, with joint block at both little fingers, where patient can't approximate palms of both hands together.

Plaques are well defined, demarcated scaly, no fissures or erythema, 1-2 cms in diameter. No hyperhidrosis is seen. Lesions are **skin colored**.

The lesions are **Firm** on palpation.

Palms & soles: Diffuse transgradient palmoplantar keratoderma. **No** Palmar or plantar pits or erythema, as well as arachnodactyly or pseudoainhum of the digits, **not** detected.

Nails: Nail dystrophy with hyperconvexity, transverse and longitudinal ridging is noticed at most of dysplastic nails.

Hair: No Abnormality Detected, with normal body and scalp hair.

Eye: **No** abnormality seen.

Oral mucosa & Lips: Periodontosis with loss of teeth-Oligodontia. **No** Leukokeratosis.

5. The Case 3

A male patient, student, 8 years old, mixed race, Gaali tribe, resident in Al-Duba, north of Sudan. The patient is complaining of dry painful fissured, non-itchy, soles of feet with difficulty on walking since age of 4 months, loss of teeth with swollen painful gum since the age of 15 years old.

Patient's general condition looks well with average intelligence. The condition started at infancy, age of 4 months with peeling of soles of feet, with gradual onset and progressive course, where thickening of both soles ensues, fissured and worse with age and focal thickening in both palms. Patient noticed inability to move some of her hand fingers since childhood. Patient has been diagnosed as a case of Psoriasis and received topical treatment in the form of PUVA with more or less satisfactory result.

Family history: Parents are first degree relatives, the youngest brother of case 2.

On examination: Patient has normal facial features with diffuse transgradient palmoplantar keratoderma, and distant psoriasiform plaques-like at both knees and elbows. Hyperkeratotic knuckle pads mainly at proximal interphalangeal joints, with joint block at both little fingers, where patient can't approximate palms of both hands together.

Plaques are well defined, demarcated scaly, no fissures or erythema, 1-2 cms in diameter. No hyperhidrosis is seen. Lesions are **skin colored**.

The lesions are **Firm** on palpation.

Palms & soles: Diffuse transgradient palmoplantar keratoderma. **No** Palmar or plantar pits or erythema, as well as arachnodactyly or pseudoainhum of the digits, **not** detected.

Nails: Nail dystrophy with hyperconvexity, transverse and longitudinal ridging is noticed at most of dysplastic nails.

Hair: No Abnormality Detected, with normal body and scalp hair.

Eye: No abnormality seen.

Oral mucosa & Lips: Periodontosis with loss of teeth-Oligodontia. **No** Leukokeratosis.

6. Discussion

● Papillon-Lefevre syndrome is a rare disease, especially in Africans.

● In our cases of discussion here, the patients are a Sudanese from northern part of Sudan. One of them associated with Toxic Goiter and has been missed as a case of Psoriasis where given PUVA therapy.

● The condition has been put in due dental care where oral medicine and periodontal departments take care of the oral manifestations. The cases diagnosed provisionally as: Papillon-Lefevre syndrome, as mutilating palmoplantar keratoderma and periodontosis, where considered as the first cases been reported in Sudan.

In **conclusion**, Papillon-Lefevre syndrome is a rare disease, with solid diagnostic major and minor features, supposes never missed if features were followed properly. In our case, oral and dental care is essential, as the patient has got satisfied cosmetically and psychologically.

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