Familial leuconychia, knuckle pads, hearing loss, and palmoplantar hyperkeratosis: an additional family with Bart-Pumphrey syndrome

Jeanette C Ramer, David B Vasily, Roger L Ladda

Abstract
A family with five members who have variable findings of leuconychia, knuckle pads, hearing loss, and palmoplantar hyperkeratosis is described. The findings in these subjects are compared with those noted in previously reported patients with Bart-Pumphrey syndrome. The range of disorders which include knuckle pads as part of the phenotype is reviewed.

Knuckle pads are uncommon but benign skin lesions which appear as rounded, fleshy, well circumscribed papules over the dorsal surface of the small joints of the hands or feet. Although first described for a medical readership by Garrod in 1904, they were noted by artists during the Renaissance who incorporated them into portraits and sculptures along with facial likenesses of their subjects. In most subjects, these cutaneous growths appear in childhood then enlarge slowly over time. They can occur in multiple family members and, in this situation, appear to follow an autosomal dominant mode of transmission. Knuckle pads can be associated with Dupuytren's contracture but are most commonly found as an isolated anomaly, almost exclusively on the hands with feet rarely involved.
In 1967, Bart and Pumphrey4 reported a six generation family in which knuckle pads were present along with congenital deafness and leuconychia. Keratoderma was also apparent in some family members. Autosomal dominant inheritance was evident in this kindred. Similarities between these patients and a child who had been reported by Schwann5 in 1963 were evident to Bart and Pumphrey.4 This child had leuconychia, sensorineural deafness, knuckle pads, and keratoderma of the palms and soles, but was the only member of his family to manifest all of these findings. Syndactyly was present in several of the other 69 members of this kindred. A third family with the same constellation of findings was described by Crosby and Vidurrizaga6 in 1976. Four subjects in two generations were affected with knuckle pads, primarily on the toes, variable leuconychia, moderate to severe sensorineural hearing loss, and keratoderma of the soles and palms. Again the pattern of transmission was consistent with autosomal dominant inheritance.

In this report, we describe a three generation family with four members who have skin abnormalities and deafness as well as one member with hearing loss only. Breast cysts and cyst-like skin lesions were present in the family and her affected mother, findings which may represent an additional manifestation of Bart-Pumphrey syndrome.

Case reports

PATIENT II-3
This 24 year old white female was evaluated at the request of her dermatologist who had noted marked hyperkeratosis of the palms and soles, especially evident along the skin creases, knuckle pads involving the dorsal aspects of the proximal interphalangeal joints of both hands, mild leuconychia, and severe sensorineural deafness. Painful cysts of the breasts, axillae, and dorsum of the right forearm had developed over the previous year. She recalled that the knuckle pads had become apparent at the age of 8 years and had enlarged slowly since that time. The hearing loss was congenital and, apparently, unchanging. Hyperkeratosis had its onset in early adolescence with slow progression especially apparent on the soles of the feet.

Examination showed normal hair and teeth. Skin texture was normal except for the hands and feet. There were multiple, darkly pigmented macules and papules from 2 to 8 mm in diameter scattered over the upper chest, between the breasts, and in both axillae. There were a few scattered fleshy papules over the lower abdomen. Breast examination disclosed multiple tender cystic regions bilaterally. There was a discrete, raised, slightly hyperpigmented lesion on the dorsum of the left wrist which measured 2 × 2-5 cm. Palpation suggested that it was fluid filled (fig 1).

The palms and soles were hyperkeratotic with accentuation of crease patterns and yellow discoloration (figs 2 and 3). Knuckle pads measuring approximately 0-75 to 1 cm in diameter were present over the dorsal aspect of the proximal interphalangeal joints of all digits of the left hand as well as the thumb, first, and fifth fingers of the right hand (figs 4 and 5). Knuckle pads were not present on the feet. Mild leuconychia was present and lunulae were absent or poorly defined while nail shape and size were normal.

Severe sensorineural hearing loss was evident but an audiogram was not performed. The patient communicated using signing.

PATIENT III-2
This 2 year old male is the son of patient II-3. He had significant hearing loss in infancy. Examination showed hyperkeratosis of the palms, especially evident over the thenar eminence and between the thumb and first finger. There were no nail changes or knuckle pads. The feet were normal.
PATIENT I-4
This patient, the mother of patient II-3, is reported by family members to have had profound hearing loss, hyperkeratosis, and knuckle pads. Leuconychia was not recalled. She was diagnosed as having breast and axillary cysts as well as a growth on the palmar surface of one wrist similar to that seen in her daughter. She had died of the effects of a brain tumour at the age of 58 years.

PATIENT II-1
This 31 year old woman was not available to be examined, but she was reported to have striking hyperkeratosis with variable expressivity.

Table 1 Characteristics of patients with Bart-Pumphrey syndrome

<table>
<thead>
<tr>
<th>Previously reported subjects*+</th>
<th>Current family</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Males</td>
<td>Females</td>
</tr>
<tr>
<td>Leuconychia</td>
<td>5</td>
<td>5</td>
</tr>
<tr>
<td>Knuckle pads</td>
<td>5</td>
<td>5</td>
</tr>
<tr>
<td>Hands</td>
<td>3</td>
<td>3</td>
</tr>
<tr>
<td>Feet</td>
<td>4</td>
<td>2</td>
</tr>
<tr>
<td>Hyperkeratosis</td>
<td>3</td>
<td>3</td>
</tr>
<tr>
<td>Hands</td>
<td>3</td>
<td>2</td>
</tr>
<tr>
<td>Hearing loss</td>
<td>4</td>
<td>4*</td>
</tr>
</tbody>
</table>

*One patient is only 2 years of age.
+Two patients had mixed conductive and sensorineural hearing loss.
isolated knuckle pads and in palmoplantar keratoderma with knuckle pads. The second subtype, most usually found in subjects with Dupuytren's contracture or Touraine syndrome, shows thickening and hyperplasia of collagen bundles in the dermis in addition to epidermal changes. The histological findings of the knuckle pads noted in Bart-Pumphrey syndrome have not been reported.

No candidate genetic loci have been elucidated for any of these disorders. With the exception of Bart-Pumphrey syndrome, all involve focal overgrowth of ectodermal structures. The reported disruption of middle ear bone formation in one subject and a mixed conductive and sensorineural hearing loss in several other patients suggests mesodermal involvement in Bart-Pumphrey syndrome.

Additional detailed information regarding type and degree of hearing impairment in these subjects is needed to confirm a consistent conductive component, as this was noted in only one family. Unfortunately, the family we report was unwilling to undergo additional quantitation of hearing or further evaluation of skin lesions.

1 Garrod AE. Concerning pads upon the finger joints and their clinical relationships. BMJ 1904;ii:8.
Familial leuconychia, knuckle pads, hearing loss, and palmoplantar hyperkeratosis: an additional family with Bart-Pumphrey syndrome.

J C Ramer, D B Vasily and R L Ladda

J Med Genet 1994 31: 68-71
doi: 10.1136/jmg.31.1.68

Updated information and services can be found at:
http://jmg.bmj.com/content/31/1/68

Email alerting service
Receive free email alerts when new articles cite this article. Sign up in the box at the top right corner of the online article.

Notes