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Acquired keratodermas (and conditions with similar presentations)

by Brooks David Kimmis, MD

Acquired keratodermas				
Diagnosis	Clinical features	Associations	Treatments	Comments
Keratoderma climactericum	Focal plantar hyperkeratosis in postmenopausal middle aged and older women Favors pressure points Can fissure and cause pain with walking Palmar involvement, if present, is mild	Likely hormone related (similar findings reported in young women after oophorectomy) Obesity Cold, dry climates Sandals and backless shoes	Low dose oral retinoid Keratolytics (urea 25-40%) Topical estradiol 0.05% ointment	
Aquagenic palmoplantar keratoderma	Thickened, white pebbly papules on palmoplantar surfaces minutes after immersion in water Onset typically in 2 nd decade	Cystic fibrosis COX-2 inhibitors (celecoxib)	Botulinum toxin injection, aluminum chloride hexahydrate +/- urea cream	Acquired, autosomal dominant, and autosomal recessive variants have been described
Circumscribed palmar or plantar hypokeratosis	Well-circumscribed circular depressed papule or plaque on the palm or sole Usually focal Often seen in women ages 40-85	Prior trauma/burn	Topical calcipotriene, topical 5-FU, cryotherapy, excision have been attempted	Unknown etiology Focal thinning of stratum corneum overlying a diminished stratum granulosum with a drop-off between affected and unaffected skin



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Paraneoplastic keratodermas				
Diagnosis	Clinical features	Associations	Treatment	Comments
Diffuse PPK	Diffuse yellow hyperkeratosis of the palmar or plantar surface	Bronchial carcinoma		
Filiform/Spiny PPK	Spine-like hyperkeratotic, punctate papules on palmoplantar surfaces	Breast, colon, and kidney cancers	As with all paraneoplastic entities, treatment is targeted at the underlying malignancy	Filiform/spiny PPK can also exist as an autosomal dominant condition usually presenting in the 1 st -3 rd decades of life, in contrast to acquired filiform PPK which tends to appear after age 50

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Paraneoplastic keratodermas				
Diagnosis	Clinical features	Associations	Treatment	Comments
Tripe palms (form of malignant acanthosis nigricans)	Velvety palmoplantar, keratotic plaques	Stomach adenocarcinoma or other GI or GU adenocarcinomas	As with all paraneoplastic entities, treatment is targeted at the underlying malignancy	May or may not have findings of acanthosis nigricans in other areas (axillae, groin, nape of neck, oral mucosa)
Bazex syndrome, AKA acrokeratosis paraneoplastica (not to be confused with inherited Bazex-Dupre-Christol syndrome)	Acral psoriasiform and violaceous plaques with similar plaques on nose and/or helices. Horizontal and longitudinal nail ridging	Pharyngeal, laryngeal, and esophageal malignancies		

Keratoderma-like conditions related to exogenous substances				
Diagnosis	Clinical features	Associations	Treatment	Comments
Arsenical keratoses	Small, punctate, hyperkeratotic papules resembling corns on palms and soles Can increase in number and progress to larger, thicker papules and plaques with spread to dorsal hands and feet	Chronic arsenic exposure (contaminated well water or occupational exposure, including ore mining, agricultural use, or medicinal applications) Lesions at risk of progression to SCC in-situ, SCC, and BCC	Oral retinoids (reduces keratoses as well as risk of arsenical BCC) Can also consider excision, cryotherapy, topical 5-FU, topical imiquimod, or photodynamic therapy Chelation for chronic arsenic exposure may or may not be of benefit	Onset of arsenical keratoses ranges from 10-30 years after onset of exposure Visceral malignancies (lung and bladder) related to arsenic tend to develop after PPKs
Acral erythema AKA hand-foot syndrome AKA palmoplantar erythrodysesthesia and hand-foot skin reaction	Presentation is variable but includes diffuse edema, redness and scale of palmoplantar surfaces, and/or hyperkeratotic callus-like plaques overlying points of pressure and friction Pain, paresthesia, tingling, soreness, inability to tolerate hot substances	Prolonged infusions of chemotherapy Diffuse redness seen with cytotoxic agents such as cytarabine, doxorubicin, 5-FU Hyperkeratotic lesions seen with tyrosine kinase inhibitors such as sorafenib or sunitinib BRAF inhibitors	Symptomatic management includes topical keratolytics, pain control, topical corticosteroids, and wound care If severe, may need dose alteration or discontinuation of the drug	Exists on a spectrum of chemotherapy-induced eruptions called toxic erythema of chemotherapy, which also includes Ara-C ears, intertriginous eruption associated with chemotherapy, epidermal dysmaturation, and eccrine squamous syringometaplasia When not lumped together, "acral erythema" is considered the more severe variant with redness and edema and seen mostly with cytotoxic agents, while "hand-foot skin reaction" is the hyperkeratotic variant seen with kinase inhibitors

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Keratoderma as a presentation of other skin conditions				
Diagnosis	Clinical features	Associations	Treatments	Comments
Psoriasis	Erythematous plaques with psoriasiform, silver scale on palmoplantar surface +/-sterile palmoplantar pustules and nail pitting, onycholysis, or oil drop spots	Psoriatic arthritis Metabolic syndrome Cardiovascular disease	Topical corticosteroids +/- under occlusion, topical retinoids, topical vitamin D analogues Systemic retinoids, methotrexate, biologics, cyclosporine (for short term therapy) Phototherapy	Look for other findings of psoriasis (involvement of scalp, umbilicus, elbows, or knees, nail pitting, oil drop spots, joint swelling) Biopsy can be helpful
Chronic hand eczematous dermatitis • Allergic contact dermatitis • Irritant contact dermatitis • Other eczematous variants (atopic dermatitis, dyshidrotic eczema, others)	Erythematous scaly plaques often involving the palmar and dorsal surface. May extend to volar wrist Fissured, hyperkeratotic plaques of digits Fissuring is a common finding and is painful	Common irritants: water, detergents, fragrances, soaps, solvents, and physical irritants Common allergens: methylisothiazolinone (wet wipes), rubber, gold, nickel	Avoidance of irritants or allergens Topical barrier creams Gloves and physical barriers Topical corticosteroids Depending on the underlying cause, systemic corticosteroids, dupilumab, phototherapy	Consider patch testing for chronic eczematous hand dermatitis Biopsy can be helpful to rule out other causes of keratoderma
Pityriasis Rubra Pilaris (PRP)	Distinctive orange-red keratoderma of palms and soles		Systemic retinoids Methotrexate Topicals (corticosteroids, retinoids, keratolytics) are useful as adjuvant therapy but not as sole form of therapy	Clinical types: • Classic adult • Atypical adult • Classic juvenile • Circumscribed juvenile • Atypical juvenile • HIV-associated Types I and III tend to be self-resolving Clinical clues favoring PRP: salmon orange perifollicular keratotic papules coalescing into plaques, islands of sparing

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Diagnosis	Clinical features	Associations	Treatments	Comments
Darier disease	Punctate hyperkeratotic papules and depressed pits +/- focal keratoderma with hyperkeratotic plaques	<p>Patients are prone to secondary skin bacterial, fungal, and viral infections. (Remember that Kaposi varicelliform eruption doesn't only apply to atopic dermatitis)</p> <p>Salivary gland obstruction</p> <p>Neuropsychiatric disorders</p> <p>Ocular complications</p>	<p>Topical and systemic retinoids</p> <p>Localized areas: excision with skin grafting, laser, or dermabrasion</p>	<p>Darier is an autosomal dominant genetic disorder, but symptoms can uncommonly begin in the 2nd decade or later.</p> <p>Family history is helpful</p> <p>Exam findings favoring Darier disease include seborrheic distribution of keratotic plaques, erythronychia, V-shaped nicking of nails, verrucous brown papules on dorsal hands and feet, and oral rugose plaques</p>

Other considerations to keep in mind:

- PPK associated with lymphedema
- Palmoplantar involvement of mycosis fungoides
- Keratoderma secondary to drugs other than chemotherapeutics (lithium, verapamil, venlafaxine)
- Mechanic's hands of dermatomyositis
- Myxedema of hypothyroidism
- Keratoderma blenorrhagicum of reactive arthritis
- Crusted scabies
- Dermatophytosis

Key:

PPK, palmoplantar keratoderma

SCC, squamous cell carcinoma

BCC, basal cell carcinoma

5-FU, 5-fluorouracil

GI, gastrointestinal

GU, genitourinary

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