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Square punch differential diagnoses

By Stephanie Saridakis, DO, Melanie Wolf, DO, and Thomas Davis, MD, FAAD

Diagnosis	Histopathologic features	Clinical features and associations	Histology image
Chronic radiation dermatitis	 Hyperkeratosis common Homogenized dermis without vacuolar change or lymphoid band (vs. LSetA) Prominent dilated ectatic vessels superficially Large, stellate "radiation" fibroblasts Loss of adnexa Radiation elastosis 	Usually limited to the irradiated area Chronic changes occur months to years after initial exposure Clinically, epidermal atrophy/fragility, telangiectasias, hypoor hyperpigmentation and alopecia are present	
Necrobiosis lipoidica	 Diffuse granulomatous inflammation with multinucleated giant cells Acellular pale degenerated collagen between the layers of granulomatous inflammation → "layered cake" appearance Dermal sclerosis (late stage) Increased plasma cells deep in dermis No mucin or eosinophils (vs. GA) +/- cholesterol clefts or lymphoid nodules 	 Violaceous to pink-brown plaques with palpable peripheral rim and telangiectasias overlying subtle yellow-brown atrophic centers Typically located on anterior shins +/- ulceration Only 0.3% of patients with diabetes have NL 	
Normal back skin	 Normal thickness of collagen bundles, extending deep within the dermis Broad fascicles of collagen Lacks prominent mucin (vs. scleredema) 		
Scleroderma/ morphea	 Thick hyalinized collagen bundles Loss of adventitial fat → "trapped" eccrine glands Sparse lymphoplasmacytic infiltrate at dermal-subcutaneous junction Reduced CD34+ interstitial cells in dermis (vs. increased in NSF) +/- superficial pallor 	Group of diseases including limited systemic sclerosis, diffuse systemic sclerosis, and localized cutaneous diseases of morphea or linear scleroderma Localized scleroderma commonly presents as asymmetric sclerotic plaques Active lesions have lilac color border	

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Diagnosis	Histopathologic features	Clinical features and associations	Histology image
Scleredema	Thick dermis Increased mucin deposition predominantly in the deep dermis with widened spaces between collagen bundles No inflammation or fibrosis	Woody induration of skin, commonly involving the upper back Associations include diabetes mellitus, streptococcal infection, and monoclonal gammopathy	
Scar	 Fibroblasts arranged horizontally in an east-west direction Blood vessels arranged vertically in a north-south direction Decreased elastic tissue Effaced epidermis Loss of adnexa 	Flesh-colored to pink, hyperpigment- ed, or hypopig- mented, indurated or atrophic plaque at the site of prior trauma	
Connective tissue nevus	Collagenoma Thick, haphazard bundles of collagen with widely spaced elastic fibers Elastoma Normal to somewhat thickened dermis Increased numbers of thick, irregular, fragmented elastic fibers in the mid-tolower dermis with elastic tissue stains	Collagenoma Skin colored cerebriform plaque Associated with Tuberous Sclerosis (AD, TSC1, or TSC2 mutations, known as shagreen patch) or Proteus Syndrome (somatic, AKT1 mutation) Can be inherited in an autosomal-dominant fashion Elastoma Flesh-colored to yellow papules or plaques Associated with Buschke-Ollendorff Syndrome (AD, LEMD3 mutation)	
Sclerodermoid chronic graft- versus-host disease	Subtle basal layer vacuolization Thick, sclerotic dermis with loss of adnexal structures Oftentimes indistinguishable from morphea/scleroderma without history	Non-sclerotic cGVHD: • Many morphologies (lichenoid, psoriasiform, poikilodermatous, atopic dermatitis-like, lupus-like, or keratosis pilarislike eruptions) Sclerotic cGVHD: • Lichen sclerosuslike, morphea/scleroderma-like, or eosinophilic fasciitislike plaques	

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Key

AD	Autosomal dominant
cGVHD	Chronic graft-versus-host disease
GA	Granuloma annulare
LSetA	Lichen sclerosus et atrophicus
NL	Necrobiosis lipoidica
NSF	Nephrogenic systemic fibrosis

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Histology slides courtesy of Tom Davis, MD, FAAD

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