Distinguishing cellulitis from its mimics

ABSTRACT

Distinguishing true cellulitis from its many imitators is challenging but critical if we are to avoid unnecessary use of antibiotics and delays in treatment. Common imitators of cellulitis are stasis dermatitis, lipodermatosclerosis, contact dermatitis, lymphedema, eosinophilic cellulitis, and papular urticaria. Specific criteria do not exist for the diagnosis of cellulitis, but the alert physician can find clues in the history and physical examination that point toward cellulitis.

KEY POINTS

Cellulitis is rarely bilateral.

Patients with cellulitis often have systemic symptoms, such as fever and leukocytosis.

A chronic course points to a diagnosis other than cellulitis.

Plaques with a “bound-down” appearance or dark pigmentation point to a chronic disease rather than cellulitis.

Stasis dermatitis is the most common mimic of cellulitis.

More than 10% of patients labeled as having cellulitis do not have cellulitis. This is unfortunate, as it leads to excessive and incorrect use of antibiotics and to delays in appropriate therapy. However, it is not surprising, given the number of conditions that bear a striking similarity to cellulitis. A familiarity with the features of true cellulitis and with the handful of conditions that can bear a striking similarity to it is the way out of this potential diagnostic quagmire.

WHAT CELLULITIS IS—AND IS NOT

The key characteristics of cellulitis are redness, warmth, tenderness, and swelling of the skin. A history of trauma and pain in the affected area and evidence of leukocytosis suggest cellulitis. A symmetric or diffusely scattered pattern indicates a condition other than cellulitis, which is overwhelmingly unilateral, with smooth, indistinct borders. Other factors pointing to cellulitis are underlying immunosuppression, a more rapid progression, previous episodes, systemic symptoms (eg, fever, leukocytosis), new medications, new travel or outdoor exposure, and comorbidities such as diabetes and peripheral vascular disease. A long-standing, slowly progressive course and a history of unsuccessful treatment with antibiotics are strong indicators of a condition other than cellulitis.

Consultation with a dermatologist is recommended to narrow the differential diagnosis. The dermatologist can determine if biopsy is necessary, as many dermatoses that mimic cellulitis can be diagnosed by visual recognition alone.
The most common mimic of cellulitis is stasis dermatitis (figure 1).

The inciting factor is chronic venous insufficiency, leading to interstitial edema, extravasation of red blood cells, and decreased tissue oxygenation. This process causes microvascular changes and microthrombi that upregulate transforming growth factor beta and fibroblastic growth factor. If the process is allowed to continue, stasis dermatitis may progress to lipodermatosclerosis.

Tip: Stasis dermatitis is generally bilateral, the process will have been ongoing for years, there is often pitting edema, and the legs should be nontender.

Lipodermatosclerosis is a sclerosing panniculitis classically described as an “inverted champagne bottle” or “inverted bowling pin” appearance of the legs, ie, the diameter of the leg is sharply narrowed directly below the calf (figure 2).

There is an acute and a chronic phase. The acute phase is characterized by inflammation and erythema, and the chronic phase is characterized by fibrosis. The acute phase presents with severe lower-extremity pain above the medial malleolus, erythema, edema, and warmth; there is no sharp demarcation between affected and unaffected skin. This phase can be difficult to distinguish from cellulitis, so the history plays a key role. Known venous insufficiency, cutaneous changes of stasis dermatitis, and the absence of systemic symptoms all point to lipodermatosclerosis.

The chronic phase is characterized by unilateral or bilateral, indurated, sclerotic plaques with a “bound-down” appearance (ie, they appear as if tethered—or bound—to the subcutaneous tissue) affecting the skin from below the knee to the ankle; there is a sharp demarcation between affected and unaffected skin. The skin is often bronze or brown secondary to hemosiderin deposits. There can be prominent varicosities and scattered ulcerations depending on the course of the disease.

This condition is thought to be the result of long-standing chronic venous insufficiency. It is proposed that venous incompetence leads
to extravasation of interstitial fluid and red blood cells, decreased diffusion of oxygen to the tissues, and eventual tissue and endothelial damage. As the endothelium is damaged, microthrombi formation and infarction ensue, stimulating fibroblasts to form granulation tissue.

Tip: The history helps to distinguish acute lipodermatosclerosis from cellulitis. Chronic lipodermatosclerosis will have been ongoing for years, the legs should be nontender, the skin will be bound-down, and the diameter of the leg will sharply decrease from knee to ankle.

**CONTACT DERMATITIS**

Allergic and irritant forms of contact dermatitis are often mistaken for cellulitis. Irritant contact dermatitis (figure 3) presents with erythematous patches and plaques with well-defined borders, often in a geometric distribution where the skin was exposed to an irritant. Allergic contact dermatitis is a delayed hypersensitivity dermatitis that can be secondary to something ingested, applied to the skin, or airborne (figure 4). It presents as erythematous macules, papules, and plaques that may have serous drainage or vesiculation. Lesions of allergic contact dermatitis are usually confined to the site of contact with the allergen, but they can infrequently be found at distant sites, in which case it is considered systemic contact dermatitis. Depending on the severity of the allergy, patients may complain of intense pain and pruritus.

Additionally, chronic, nonhealing leg ulcers may have a confounding allergic contact dermatitis. Although patients may believe they are helping the ulcer heal by applying topical antibiotics or other lubricants, they may in fact be impeding the healing process. Always inquire as to what the patient is applying if he or she has leg ulceration with surrounding edema and erythema that has not resolved with conventional treatments.

Tip: The key to distinguishing contact dermatitis from cellulitis is the history. For example, ask about recent changes in medications, soaps, and laundry detergents, new hobbies, or recent surgeries. The involved site is often confined to the area where the allergen contacted the skin, except in cases of exposure to an airborne allergen.

**LYMPHEDEMA**

Lymphedema is characterized by localized edema of an affected extremity, with induration, erythema, and secondary cutaneous changes such as hyperkeratosis, dyspigmentation, and wart-like architecture (figure 5). Primary lymphedema appears in the setting of congenital abnormalities, whereas secondary lymphedema results from an interruption of a previously functioning lymphatic system (eg, after radical mastectomy).

Patients often present with unilateral non-pitting edema and erythema in the absence of systemic symptoms. Many patients presenting with lower-extremity lymphedema are overweight or obese, as the weight they carry causes obstruction of the inguinal lymphatics.
The pathophysiology is not clearly delineated but is thought to be a consequence of decreased oxygenation of tissue secondary to extravasated lymph. As the oxygen is compromised, macrophages and fibroblasts are recruited, resulting in fibrosis.\(^6\)

Patients with lymphedema are more susceptible to superficial and deep skin infections, as the natural defense system in the epidermis and papillary dermis is compromised by impaired lymphatic drainage.\(^15\)

To differentiate uncomplicated lymphedema from a secondary cutaneous infection, the clinician should take into account the presence or absence of warmth, pain, increased erythema, and systemic symptoms (FIGURE 6).

**Tip:** Primary lymphedema will most likely present in childhood with no inciting factors and will require a full workup. Obtaining a history should make secondary lymphedema a relatively straightforward diagnosis: Has the patient undergone lymph node dissection? Has the patient had an injury in the affected leg? Lymphedema is overwhelmingly unilateral and nonpitting, and is often seen in overweight people (if no precipitating factor is present).

### EOSINOPHILIC CELLULITIS

Eosinophilic cellulitis, or Wells syndrome, was first described in 1971 as a granulomatous dermatitis.\(^16\) It is a recurrent hypersensitivity reaction to a drug, to a vaccine, or to an insect bite, or to a viral or fungal infection that presents on the extremities as localized erythema, edema, and induration with sharp borders and a green or gray hue (FIGURE 7).\(^17,19\) The lesions commonly progress to firm, indurated plaques that resemble morphea. The plaques may take weeks or years to resolve, but they do so without scarring.\(^12,17,20,21\)

**Tip:** This diagnosis often requires biopsy for confirmation, but helpful clues are a history of recurrent episodes, the color of the lesions, and peripheral eosinophilia.

### PAPULAR URTICARIA

Papular urticaria is a dermal hypersensitivity reaction to an insect bite, most commonly from a flea or mosquito.\(^26\) Patients are often children, as their immune system may be hypersensitive. But children often develop tolerance before puberty.\(^27\)
The presentation may vary, from numerous urticarial papules near the site of a bite, to generalized, large, indurated, erythematous plaques reminiscent of cellulitis (FIGURE 8).5,26 The lesions usually develop within hours of a bite and persist for an average of 1 to 2 weeks.28 The areas typically affected are the head and neck or the upper or lower extremities; the palms, soles, and trunk are usually spared.27

Patients most often complain of intense itching.12 The pathogenesis is proposed to be mediated by the immune complex, and tissue biopsy study shows increased eosinophils. The eosinophils stimulate mast cells, causing release of histamine, leading to increased vascular permeability, edema, and erythema.28,29

**Tip:** Biopsy may be necessary to confirm the diagnosis, though often the history may be sufficient. The patient may or may not recall a bite, so probe into recent activities such as outdoor sports or contact with a new pet. The papules and plaques are generally very pruritic but not painful.

**DERMATOLOGY CONSULT**

If the clinical presentation and history do not correlate, or if the skin condition has been treated with antibiotics yet has failed to respond, the possibility of other cutaneous dermatoses should be entertained. A dermatology consult can help determine the diagnosis, the need for further evaluation, and the best treatment course.

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**REFERENCES**

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