

Acneiform facial eruptions

A problem for young women

Melody J. Cheung, MD Muba Taher, MD, FRCPC Gilles J. Lauzon, MD, PHD, FRCPC

ABSTRACT

OBJECTIVE To summarize clinical recognition and current management strategies for four types of acneiform facial eruptions common in young women: acne vulgaris, rosacea, folliculitis, and perioral dermatitis.

QUALITY OF EVIDENCE Many randomized controlled trials (level I evidence) have studied treatments for acne vulgaris over the years. Treatment recommendations for rosacea, folliculitis, and perioral dermatitis are based predominantly on comparison and open-label studies (level II evidence) as well as expert opinion and consensus statements (level III evidence).

MAIN MESSAGE Young women with acneiform facial eruptions often present in primary care. Differentiating between morphologically similar conditions is often difficult. Accurate diagnosis is important because treatment approaches are different for each disease.

CONCLUSION Careful visual assessment with an appreciation for subtle morphologic differences and associated clinical factors will help with diagnosis of these common acneiform facial eruptions and lead to appropriate management.

RÉSUMÉ

OBJECTIF Faire le point sur le diagnostic clinique et les modalités thérapeutiques actuelles de quatre types d'éruptions faciales acnéiformes chez la femme jeune: l'acné vulgaire, l'acné rosacée, la folliculite et la dermatite périorale.

QUALITÉ DES PREUVES Le traitement de l'acné vulgaire a fait l'objet de plusieurs essais randomisés ces dernières années. Les recommandations pour le traitement de l'acné rosacée, de la folliculite et de la dermatite périorale reposent surtout sur des essais comparatifs ou ouverts (preuves de niveau II), mais aussi sur des opinions d'experts et des déclarations de consensus (preuves de niveau III).

PRINCIPAL MESSAGE Les femmes jeunes consultent fréquemment les établissements de soins primaires pour des éruptions faciales acnéiformes. Il est souvent difficile de distinguer des conditions morphologiquement semblables. Il importe toutefois de poser un diagnostic précis car les modalités thérapeutiques diffèrent d'une maladie à l'autre.

CONCLUSION Le diagnostic et le traitement des éruptions faciales communes sont plus faciles si l'on fait une évaluation visuelle attentive et si on tient compte des différences morphologiques subtiles et des facteurs cliniques associés.

This article has been peer reviewed. Cet article a fait l'objet d'une évaluation externe.



cneiform eruptions, such as acne vulgaris, rosacea, folliculitis, and perioral dermatitis, are routinely encountered in primary care. Acne vulgaris alone affects up to 80% of adolescents and continues to affect 40% to 50% of adult women.¹ An estimated 13 million Americans are affected by rosacea.² These conditions often have psychosocial sequelae.³

These four eruptions are challenging to diagnose because they all resemble acne. This article describes these eruptions, highlighting the salient distinguishing characteristics, and summarizes current management recommendations from the medical literature.

Quality of evidence

PubMed was searched from January 1966 to December 2003 using the names of each of the acneiform conditions combined with "treatment." Several randomized controlled trials (level I evidence) on treatment of acne vulgaris were found, but there was little level I evidence for treating the other conditions. Recommendations for treating these conditions are based mainly on comparison or open-label studies (level II evidence) and expert opinion and consensus guidelines (level III evidence).

Acne vulgaris

Acne vulgaris is a disease of the sebaceous follicles that primarily affects adolescents but not uncommonly persists through the third decade and beyond, particularly in women. Pathogenesis is multifactorial and involves an interplay between abnormal follicular keratinization or desquamation, excessive sebum production, proliferation of follicular *Propionibacterium acnes*, and hormonal factors.

Diagnosis is often clear, and laboratory investigations are unnecessary, except where signs and symptoms suggest hyperandrogenism.^{4,5} Acne is

Dr Cheung is a dermatology resident, Dr Taher has completed dermatology residency, and Dr Lauzon is an Associate Professor and Director in the Division of Dermatology, all at the University of Alberta in characterized by a variety of lesions that indicate varying degrees of disease severity.

Mild or noninflammatory acne is characterized by comedones. Closed comedones appear as pale white, slightly elevated, dome-shaped, 1- to 2-mm papules with no clinically visible follicular orifice (Figure 1). Open comedones are flat or slightly raised lesions with a visible central orifice filled with a brown-black substance (Figure 1). Inflammatory acne has a range of lesions. Papules (Figure 1) are often encircled by an inflammatory halo, and pustules can be identified by a central core of purulent material. Nodules are rounder and deeper to palpation than papules and are often tender. Cysts have a propensity to scar and essentially feel like fluctuant nodules. Acne scars (Figure 1) usually appear as sharply punched out pits.

Figure 1. Acne vulgaris in various stages: A) Several closed comedones (1-mm to 2-mm pale white, dome-shaped papules); B) Several open comedones, papules with a central orifice filled with a brown-black substance; C) Acne papule; D) Several punched-out depressions marking acne scars.



Before commencing therapy and in the interest of establishing a therapeutic alliance, it is important to explain to patients the causes of acne and the rationale for therapy as well as the expected duration of therapy (weeks to months). The literature suggests that therapy be based on the severity or the predominant morphologic variant of disease.

Mild comedonal acne should be treated with topical antimicrobials, 1.6-8 such as benzoyl perox-



topical comedolytics,1,6-8 such as tretinoin (available in 0.025/0.05/0.1% cream, 0.01/0.025% gel, and 0.05% liquid) (Table 19-24). Benzoyl peroxide is preferred for patients with inflammation8; tretinoin is effective for cases with a predominance of comedones. The recently developed topical retinoid, adapalene, is only marginally more effective than tretinoin, but is better tolerated.7 Choice of treatment depends largely on patients' tolerance and preference.6 Gels and creams with water bases are less drying than gels in alcohol or glycol bases. Exfoliants, such as salicylic acid, remain an option for acne treatment, but are ineffective for deep comedones and can be irritating.1

Papular and pustular acne can be treated with topical or oral antibiotics (Table 19-24). Both topical erythromycin (available as solution, gel, or pledgets) and clindamycin (available as solution, gel, lotion, or pledgets) are reported to be equally effective. 9,25 Topical erythromycin is considered safest during pregnancy.1

Combination topical products, such as Clindoxyl (clindamycin and benzoyl peroxide) and Benzamycin (benzoyl peroxide and erythromycin), have recently come on the market and are quite useful.6 Tetracycline (1000 mg/d in two or four divided doses), because of its effectiveness and low cost, is the first-choice oral antibiotic followed by minocycline (50 to 100 mg/d) or doxycycline (100 mg/d). These drugs are often prescribed, along with topical retinoids, combination products, or antimicrobials, to improve efficacy and prevent resistance from developing. Trimethoprim-sulfamethoxazole is best reserved for severe, recalcitrant cases.1 Other oral antibiotics mentioned in the literature include erythromycin, clindamycin, ampicillin, and amoxicillin in no particular order. Most of these drugs should be used for at least 2 months before they are deemed ineffective.6

Cases of treatment-resistant, nodulocystic, or scarring acne should be referred to a dermatologist for isotretinoin treatment, steroid injection, or hormone therapy (Table 19-24). Isotretinoin is notorious for its drying side effects and teratogenicity, but is a very effective medication with a response rate as high as 90%.1 It is administered at 0.5 to 1.0

Table 1. Recommendations for treating acne vulgaris

NONINFLAMMATORY

First line: Benzoyl peroxide (I)10 or topical tretinoin (I)11 Second line: Adapalene (I)¹¹

PAPULAR OR PUSTULAR

First line: Topical erythromycin (I), 10,12 clindamycin (I), 9,13 clindoxyl (I), 14 or benzamycin (I)15

Second line: Oral tetracycline (I),16 minocycline (I),17 doxycycline (I),18 erythromycin (I),16 clindamycin (I),19 ampicillin (III), or amoxicillin (III) Third line: Oral antibiotics plus topical retinoids (I),20 clindoxyl or benzamycin (III), or antimicrobials (III) Fourth line: Trimethoprim-sulfamethoxazole (III)

NODULOCYSTIC OR TREATMENT-RESISTANT ACNE OR SCARRING

First line: Steroid injection, if sparse (I), 21 isotretinoin, if diffuse (I)22 Second line: Antiandrogens, for example, oral contraceptives (I)²³ or spironolactone (I)24

Roman numerals indicate level of evidence.

early response with minimal side effects. Average duration of therapy is 4 months; a second course might be necessary. Triamcinolone acetonide intralesional injections are feasible for sparser nodulocystic lesions, but care must be taken to avoid steroid atrophy. Finally, for women unresponsive to conventional therapy, hormonal therapy (biphasic or triphasic contraceptive pills or spironolactone, which has strong antiandrogenic activity) is recommended in conjunction with topical treatment.^{1,8}

Rosacea

Rosacea is a chronic vascular acneiform facial disorder that affects primarily 20- to 60-year-old people of northern and eastern European descent. Although the condition is equally prevalent in men and women, it is usually more severe in men and can progress to tissue hyperplasia. Pathogenesis remains unknown, although many factors including bacteria, Demodex mites, vasomotor and connective tissue dysfunction, and topical corticosteroids have been implicated.

Rosacea is characterized by a triad of symmetrical erythema, papules and pustules, and telangiectasia on the cheeks, forehead, and nose (Figures 2 and 3). The absence of comedones is an important factor that differentiates rosacea from acne vulgaris. Rosacea follows a course of exacerbations and



Figure 2. Rosacea: A young woman has persistent erythema and red



Figure 3. Rosacea: Rosacea can persist into later decades. This older woman has deep symmetrical erythema and many red papules on her cheeks, forehead, and chin.



Table 2. Recommendations for treating rosacea

First line: Topical metronidazole (I)³⁰ plus oral tetracycline (I)³¹ or minocycline (III) Second line: Sulfacetamide (III) plus oral antibiotics as above Third line: Topical retinoid (II)³² plus vitamin C (II)²⁹ Fourth line: Isotretinoin (II)³³

Roman numerals indicate level of evidence.

and hot drinks. Frequent flushing, mild telangiectasia, increased telangiectasia with acneiform eruptions, and tissue hyperplasia are the four sequential stages of the condition. Rosacea can also be associated with ocular symptoms of burning, redness, itching, sensation of a foreign body, tearing, dryness, photophobia, and eyelid fullness or swelling.²⁶

Begin treatment by discussing potential triggers and how to avoid them. Concomitant topical metronidazole and oral tetracycline are recommended as first-line therapy for early-stage rosacea^{27,28} (Table 2²⁹⁻³³). This combination lowers the potential for relapse once the oral medication is withdrawn.^{27,28} Oral minocycline (100 to 200 mg/d) is considered an acceptable alternative.²⁸ Doxycycline, clindamycin, erythromycin, clarithromycin, ampicillin, and metronidazole have also been shown to be effective (Table 2²⁹⁻³³). Oral therapy should be prolonged in those with ocular symptoms, although some sources recommend deferring oral antibiotics until there are ocular complaints.³⁴

There is no significant difference in efficacy between twice-daily treatment with 0.75% topical metronidazole and once-daily treatment with 1.0% metronidazole.27 Topical sulfacetamide is an alternative if metronidazole is not tolerated or if patients want concealment (sulfacetamide is available in a flesh-coloured preparation) (Table 2²⁹⁻³³). Oral tetracycline is usually started at 1000 mg/d, tapered, and finally discontinued. Various sources recommend various tapering protocols and duration of therapy. Some recommend tapering to 500 mg/d over 6 weeks followed by a slow maintenance taper to 250 mg/d over 3 months if patients respond; otherwise, a 6-week course of full-dose tetracycline should be repeated.2 Others recommend therapy at full dose until clearance or for 12 weeks' duration.²⁸ Recently, topical retinoid and vitamin C preparations have been shown to have a beneficial effect^{29,32} (Table 2²⁹⁻³³).

For recalcitrant rosacea, a 4- to 5-month course of oral isotretinoin at either low dose (10 mg/d) or the dose used for acne vulgaris has been shown to reduce symptoms.³⁵ Patients with rosacea with fibrotic changes should be referred to a cosmetic



Folliculitis

Folliculitis is an inflammation of the hair follicle as a result of mechanical trauma (eg, shaving, friction), irritation (certain topical agents, such as oils), or infection. Mechanical trauma, occlusion, and immunocompromise predispose patients to infection. The usual infectious organism is Staphylococcus aureus, although Gram-negative folliculitis can result from prolonged use of antibiotics for acne. Pityrosporum, a saprophytic yeast, has also been implicated.

Diagnosis is clinical. There is usually an abrupt eruption of small, well circumscribed, globular, dome-shaped, often monomorphic pustules in clusters on hair-bearing areas of the body and face (Figure 4). Deeper follicular infections, or sycosis, although rare, are more erythematous and painful.

Initially, potassium hydroxide testing of the hair and any surrounding scale should be considered to exclude Pityrosporum. Otherwise, an identifying culture should always be taken before initiating therapy.³⁴ In confirmed cases, topical therapy with econazole cream, selenium sulfide shampoo, or 50% propylene glycol³⁶ has been recommended for a duration of 3 to 4 weeks (Table 337-41). Subsequent additional

Figure 4. Folliculitis: A cluster of small monomorphic pustules appears on a woman's forehead.

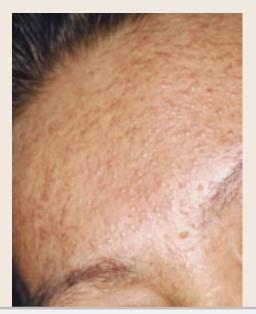


Table 3. Recommendations for treating folliculitis

PITYROSPORUM

First line: Topical econazole (III), selenium sulfide shampoo (III), or 50% propylene glycol (III)

Second line: Oral fluconazole (II),³⁷ itraconazole (I),³⁸ or ketoconazole (II)³⁷

Third line: Oral antifungal plus topical agents (II)³⁷

BACTERIAL

First line: Topical mupirocin (I), ³⁹ erythromycin (III), clindamycin (III), or benzoyl peroxide (III)

Second line: Oral antistaphylococcal antibiotics, such as fluoroquinolones (I),40 first-generation cephalosporins (III), or macrolides (III)

GRAM NEGATIVE

First line: Isotretinoin (II)41

Second line: Ampicillin (III) or trimethoprim-sulfamethoxazole (III)

Roman numerals indicate level of evidence.

intermittent maintenance doses once to twice a week42 have been found helpful for avoiding recurrence, which is common in folliculitis. Oral antifungals (fluconazole, ketoconazole, or itraconanzole) have been deemed effective when used for 10 to 14 days⁴³ (Table 3³⁷⁻⁴¹). One clinical trial demonstrated the superiority of combined topical and oral therapy as compared with either alone.37

Topical therapy for superficial *S aureus* includes erythromycin, clindamycin, mupirocin, or benzoyl peroxide44 (Table 337-41). Oral antistaphylococcal antibiotics (first-generation cephalosporins, penicillinase-resistant penicillins, macrolides, or fluoroquinolones) are indicated for extensive disease or for the deep involvement of sycosis44 (Table 3³⁷⁻⁴¹). Treatment is continued until lesions completely resolve. 45 Gram-negative folliculitis can be treated as severe acne with isotretinoin at a dose of 0.5 to 1.0 mg/kg daily for 4 to 5 months⁴⁶ (Table 337-41). Alternatives are ampicillin at 250 mg or trimethoprim-sulfamethoxazole at 600 mg four times daily, but response to antibiotic treatment is slow, and relapse is common.

Perioral dermatitis

Perioral dermatitis is an acneiform eruption of unknown etiology, although many contribut-



DOCKET

Explore Litigation Insights



Docket Alarm provides insights to develop a more informed litigation strategy and the peace of mind of knowing you're on top of things.

Real-Time Litigation Alerts



Keep your litigation team up-to-date with **real-time** alerts and advanced team management tools built for the enterprise, all while greatly reducing PACER spend.

Our comprehensive service means we can handle Federal, State, and Administrative courts across the country.

Advanced Docket Research



With over 230 million records, Docket Alarm's cloud-native docket research platform finds what other services can't. Coverage includes Federal, State, plus PTAB, TTAB, ITC and NLRB decisions, all in one place.

Identify arguments that have been successful in the past with full text, pinpoint searching. Link to case law cited within any court document via Fastcase.

Analytics At Your Fingertips



Learn what happened the last time a particular judge, opposing counsel or company faced cases similar to yours.

Advanced out-of-the-box PTAB and TTAB analytics are always at your fingertips.

API

Docket Alarm offers a powerful API (application programming interface) to developers that want to integrate case filings into their apps.

LAW FIRMS

Build custom dashboards for your attorneys and clients with live data direct from the court.

Automate many repetitive legal tasks like conflict checks, document management, and marketing.

FINANCIAL INSTITUTIONS

Litigation and bankruptcy checks for companies and debtors.

E-DISCOVERY AND LEGAL VENDORS

Sync your system to PACER to automate legal marketing.

