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The many variants of psoriasis

Erythema, skin thickening, and scales typify most cases, but other patterns exist. Tx choices may differ, depending on how much body surface area is covered.

PRACTICE RECOMMENDATIONS

- Consider guttate psoriasis if small (often < 1 cm) pink scaly papules appear suddenly, particularly in a child who has an upper respiratory tract infection.
- Document extent of disease using a tool such as the Psoriasis Area and Severity Index, which calculates a score based on the area (extent) of involvement surrounding 4 major anatomical regions. ©
- Consider prescribing
 UV light treatment or a
 combination of alcitretin and
 topical corticosteroid if
 > 10% of the body surface area
 is involved but joints are not
 affected. ©

Strength of recommendation (SOR)

- Good-quality patient-oriented evidence
- B Inconsistent or limited-quality patient-oriented evidence
- Consensus, usual practice, opinion, disease-oriented evidence, case series

he "heartbreak of psoriasis," coined by an advertiser in the 1960s, conveyed the notion that this disease was a cosmetic disorder mainly limited to skin involvement. John Updike's article in the September 1985 issue of *The New Yorker*, "At War With My Skin," detailed Mr. Updike's feelings of isolation and stress related to his condition, helping to reframe the popular concept of psoriasis.¹ Updike's eloquent account describing his struggles to find effective treatment increased public awareness about psoriasis, which in fact affects other body systems as well.

The overall prevalence of psoriasis is 1.5% to 3.1% in the United States and United Kingdom.^{2,3} More than 6.5 million adults in the United States > 20 years of age are affected.³ The most commonly affected demographic group is non-Hispanic Caucasians.

Our expanding knowledge of pathogenesis

Studies of genetic linkage have identified genes and single nucleotide polymorphisms associated with psoriasis.⁴ The interaction between environmental triggers and the innate and adaptive immune systems leads to keratinocyte hyperproliferation. Tumor necrosis factor (TNF), interleukin (IL) 23, and IL-17 are important cytokines associated with psoriatic inflammation.⁴ There are common pathways of inflammation in both psoriasis and cardiovascular disease resulting in oxidative stress and endothelial cell dysfunction.⁴ Ninety percent of early-onset psoriasis is associated with human leukocyte antigen (HLA)-Cw6.⁴ And alterations in the microbiome of the skin may contribute, as reduced microbial diversity has been found in psoriatic lesions.⁵

Comorbidities are common

Psoriasis is an independent risk factor for diabetes and major adverse cardiovascular events.⁶ Hypertension, dyslipidemia, inflammatory bowel disease, nonalcoholic fatty liver disease, chronic kidney disease, and lymphoma (particularly cutane-

ous T-cell lymphoma) are also associated with psoriasis. Psoriatic arthritis is frequently encountered with cutaneous psoriasis; however, it is often not recognized until late in the disease course.

There also appears to be an association among psoriasis, dietary factors, and celiac disease. Positive testing for IgA anti-endomysial antibodies and IgA tissue transglutaminase antibodies should prompt consideration of starting a gluten-free diet, which has been shown to improve psoriatic lesions. In addition to its impact on physical health, cutaneous psoriasis often affects mental health. Increased anxiety, depression, and sleep disorders are commonly encountered, revealing the far-reaching effects of psoriasis. The persistent associated itch of psoriasis is often distressing and negatively impacts the patient's quality of life.

The different types of psoriasis

The classic presentation of psoriasis involves stubborn plaques with silvery scale on extensor surfaces such as the elbows and knees. The severity of the disease corresponds with the amount of body surface area affected. While plaque-type psoriasis is the most common form, other patterns exist. Individuals may exhibit 1 dominant pattern or multiple psoriatic variants simultaneously. Most types of psoriasis have 3 characteristic features: erythema, skin thickening, and scales.

Certain history and physical clues can aid in diagnosing psoriasis; these include the Koebner phenomenon, the Auspitz sign, and the Woronoff ring. The Koebner phenomenon refers to the development of psoriatic lesions in an area of trauma (FIGURE 1), frequently resulting in a linear streak-like appearance. The Auspitz sign describes the pinpoint bleeding that may be encountered with the removal of a psoriatic plaque. The Woronoff ring is a pale blanching ring that may surround a psoriatic lesion.

■ Chronic plaque-type psoriasis (FIGURES 2A and 2B), the most common variant, is characterized by sharply demarcated pink papules and plaques with a silvery scale in a symmetric distribution on the extensor surfaces, scalp, trunk, and lumbosacral areas.

FIGURE 1 Psoriasis appearing on traumatized tissue



Koebner phenomenon refers to psoriatic lesions that may occur in an area of trauma. Lesions may also have a linear streak-like appearance (around shirt and leg of tattoo).

■ Guttate psoriasis (FIGURE 3) features small (often < 1 cm) pink scaly papules that appear suddenly. It is more commonly seen in children and is usually preceded by an upper respiratory tract infection, often with *Streptococcus*. ¹⁰ If strep testing is positive, guttate psoriasis may improve after appropriate antibiotic treatment.

■ Erythrodermic psoriasis (FIGURES 4A and 4B) involves at least 75% of the body with erythema and scaling.¹¹ Erythroderma can be caused by many other conditions such as atopic dermatitis, a drug reaction, Sezary syndrome, seborrheic dermatitis, and pityriasis rubra pilaris. Treatments for other conditions in the differential diagnosis can potentially make psoriasis worse. Unfortunately, findings on a skin biopsy are often nonspecific,



FIGURE 2 Plaque psoriasis





Sharply demarcated pink plaques with silvery scales are commonly seen on the extensor surfaces (A), the scalp (B), trunk, and lumbosacral area.

FIGURE 3
Guttate psoriasis



The small, pink scaly papules of guttate psoriasis appear suddenly, and usually in children who have an upper respiratory tract infection.

making careful clinical observation crucial to arriving at an accurate diagnosis.

- Pustular psoriasis is characterized by bright erythema and sterile pustules. Pustular psoriasis can be triggered by pregnancy, sudden tapering of corticosteroids, hypocalcemia, and infection. Involvement of the palms and soles with severe desquamation can drastically impact daily functioning and quality of life.
- Inverse or flexural psoriasis (FIGURES 5A and 5B) is characterized by shiny, pink-to-red sharply demarcated plaques involving intertriginous areas, typically the groin, inguinal crease, axilla, inframammary regions, and intergluteal cleft.
 - **Geographic tongue** describes psoria-

FIGURE 4
Erythrodermic psoriasis





This form of psoriasis involves at least 75% of the body with erythema and scaling, and can be difficult to differentiate from such conditions as atopic or seborrheic dermatitis and pityriasis rubra pilaris.

FIGURE 5
Inverse psoriasis





Shiny, pink-to-red sharply demarcated plaques involve intertriginous areas such as the groin (A) and intergluteal cleft (B).

sis of the tongue. The mucosa of the tongue has white plaques with a geographic border. Instead of scale, the moisture on the tongue causes areas of hyperkeratosis that appear white.

- Nail psoriasis can manifest as nail pitting (FIGURE 6), oil staining, onycholysis (distal lifting of the nail), and subungual hyperkeratosis. Nail psoriasis is often quite distressing for patients and can be difficult to treat.
- Palmoplantar psoriasis (FIGURES 7A and 7B) can be painful due to the involvement of the palms of the hands and soles of the feet. Lesions will either be similar to other psoriatic plaques with well-demarcated erythema-

FIGURE 6
Nail psoriasis



Pitting of the nails with psoriasis. Onycholysis and subungual hyperkeratosis may also occur.

tous scaling lesions or involve thickening and scale without associated erythema.

Psoriatic arthritis can cause significant joint damage and disability. Most affected individuals with psoriatic arthritis have a history of preceding skin disease. ¹² There are no specific lab tests for psoriasis; radiologic studies can show bulky syndesmophytes, central and marginal erosions, and periostitis. Patterns of joint involvement are variable. Psoriatic arthritis is more likely to affect the distal interphalangeal joints than rheumatoid arthritis and is more likely to affect the metacarpophalangeal joints than osteoarthritis. ¹³

Psoriatic arthritis often progresses insidiously and is commonly described as causing discomfort rather than acute pain. Enthesitis, inflammation at the site where tendons or ligaments insert into the bone, is often present. Joint destruction may lead to the telescoping "opera glass" digit (FIGURE 8).

■ Drug-provoked psoriasis is divided into 2 groups: drug-induced and drug-

Nail psoriasis is often quite distressing for patients and can be difficult to treat.



FIGURE 7
Palmoplantar psoriasis





Palms of the hands (A) or soles of the feet (B) exhibit psoriatic plaques with well-demarcated erythematous scaling lesions. Palmoplantar psoriasis can also involve thickening and scale without associated erythema (not shown).

FIGURE 8 Psoriatic joint damage of hand



Pictured here is arthritis mutilans of the hand. Note the shortening of the digits due to bony destruction. The resulting excess skin becomes folded and retracted in an appearance similar to telescoping "opera glasses."

aggravated. Drug-induced psoriasis will improve after discontinuation of the causative drug and tends to occur in patients without a personal or family history of psoriasis. Drugaggravated psoriasis continues to progress after the discontinuation of the offending drug and is more often seen in patients with a history of psoriasis. ¹⁴ Drugs that most commonly provoke psoriasis are beta-blockers, lithium,

and antimalarials.¹⁰ Other potentially aggravating agents include antibiotics, digoxin, and nonsteroidal anti-inflammatory drugs.¹⁰

Consider these skin disorders in the differential diagnosis

The diagnosis of psoriasis is usually clinical, and a skin biopsy is rarely needed. However, a range of other skin disorders should be kept in mind when considering the differential diagnosis.

- Mycosis fungoides is a type of cutaneous T-cell lymphoma that forms erythematous plaques that may show wrinkling and epidermal atrophy in sun-protected sites. Onset usually occurs among the elderly.
- Pityriasis rubra pilaris is characterized by salmon-colored patches that may have small areas of normal skin ("islands of sparing"), hyperkeratotic follicular papules, and hyperkeratosis of the palms and soles.
- **Seborrheic dermatitis,** dandruff of the skin, usually involves the scalp and nasolabial areas and the T-zone of the face.
- Lichen planus usually appears slightly more purple than psoriasis and typically involves the mouth, flexural surfaces of the wrists, genitals, and ankles.

• Other conditions in the differential include pityriasis lichenoides chronica, which may be identified on skin biopsy. Inverse psoriasis can be difficult to differentiate from candida intertrigo, erythrasma, or tinea cruris.

A potassium hydroxide (KOH) preparation can help differentiate psoriasis from candida or tinea. In psoriasis, a KOH test will be negative for fungal elements. Mycology culture on skin scrapings may be performed to rule out fungal infection. Erythrasma may exhibit a coral red appearance under Wood lamp examination.

If a lesion fails to respond to appropriate treatment, a careful drug history and biopsy can help clarify the diagnosis.

Document disease

It's important to thoroughly document the extent and severity of the psoriasis and to monitor the impact of treatment. The Psoriasis Area and Severity Index is a commonly used method that calculates a score based on the area (extent) of involvement surrounding 4 major anatomical regions (head, upper extremities, trunk, and lower extremities), as well as the degree of erythema, induration, and scaling of lesions. The average redness, thickness, and scaling are graded on a scale of 0 to 4 and the extent of involvement is calculated to form a total numerical score ranging from 0 (no disease) to 72 (maximal disease).

Many options in the treatment arsenal

Many treatments can improve psoriasis. 9,15-19 Most affected individuals discover that emollients and exposure to natural sunlight can be effective, as are soothing baths (balneotherapy) or topical coal tar application. More persistent disease requires prescription therapy. Individualize therapy according to the severity of disease, location of the lesions, involvement of joints, and comorbidities (FIGURE 9).¹⁵

■ If \leq 10% of the body surface area is involved, treatment options generally are explored in a stepwise progression from safest and most affordable to more involved therapies as needed: moisturization and avoidance of repetitive trauma, topical corticosteroids

(TCS), vitamin D analogs, topical calcineurin inhibitors, and vitamin A creams. Recalcitrant disease will likely require ultraviolet (UV) light treatment or a systemic agent.¹⁵

■ If > 10% of the body surface area is involved, but joints are not involved, consider UV light treatment or a combination of alcitretin and TCS. If the joints are involved, likely initial options would be methotrexate, cyclosporine, or TNF- α inhibitor. Additional options to consider are anti-IL-17 or anti-IL-23 agents. ¹⁵

If there's joint involvement. In individuals with mild peripheral arthritis involving fewer than 4 joints without evidence of joint damage on imaging, nonsteroidal anti-inflammatory drugs are the mainstay of treatment. If the peripheral arthritis persists, or if it is associated with moderate-to-severe erosions or with substantial functional limitations, initiate treatment with a conventional disease-modifying antirheumatic drug. If the disease remains active, consider biologic agents.

Case studies

Mild-to-moderate psoriasis

Patient A is a 19-year-old woman presenting for evaluation of a persistently dry, flaking scalp. She has had the itchy scalp for years, as well as several small "patches" across her elbows, legs, knees, and abdomen. Overthe-counter emollients have not helped. The patient also says she has had brittle nails on several of her fingers, which she keeps covered with thick polish.

The condition exemplified by Patient A can typically be managed with topical products.

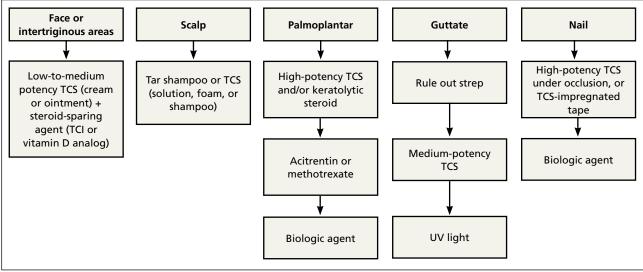
■Topical steroids may be classified by different delivery vehicles, active ingredients, and potencies. The National Psoriasis Foundation's Topical Steroids Potency Chart can provide guidance (visit www.psoriasis.org/about-psoriasis/treatments/topicals/steroids/potency-chart and scroll down). Prescribing an appropriate amount is important; the standard 30-g prescription tube is generally required to cover the entire skin surface. Ointments have a greasy consistency (typically a petroleum base), which enhances

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While plaquetype psoriasis is the most common form, other patterns exist and may even occur simultaneously.



FIGURE 9
Treatment for localized psoriasis without joint involvement¹⁵



TCI, topical calcineurin inhibitor; TCS, topical corticosteroid; UV, ultraviolet.

potency and hydrates the skin. Creams and lotions are easier to rub on and spread. Gels are alcohol based and readily absorbed. ¹⁶ Solutions, foams, and shampoos are particularly useful to treat psoriasis in hairy areas such as the scalp.

Corticosteroid potency ranges from Class I to Class VII, with the former being the most potent. While TCS products are typically effective with minimal systemic absorption, it is important to counsel patients on the risk of skin atrophy, impaired wound healing, and skin pigmentation changes with chronic use. With nail psoriasis, a potent topical steroid (including flurandrenolide [Cordran] tape) applied to the proximal nail fold has shown benefit.²⁰

- Topical calcineurin inhibitors (TCIs; eg, tacrolimus ointment and pimecrolimus cream) are anti-inflammatory agents often used in conjunction with topical steroids to minimize steroid use and associated adverse effects. ¹⁵ A possible steroid-sparing regimen includes using a TCI Monday through Friday and a topical steroid on the weekend.
- **Topical vitamin D analogs** (calcipotriene, calcipotriol, calcitriol) inhibit proliferation of keratinocytes and decrease the production of inflammatory mediators. ^{15,17-19,21} Application of a vitamin D ana-

log in combination with a high-potency TCS, systemic treatment, or phototherapy can provide greater efficacy, a more rapid onset of action, and less irritation than can the vitamin D analog used alone.²¹ If used in combination with UV light, apply topical vitamin D after the light therapy to prevent degradation.

I UV light therapy is often used in cases refractory to topical therapy. Patients are typically prescribed 2 to 3 treatments per week with narrowband UVB (311-313 nm), the excimer laser (308 nm), or, less commonly, PUVA (UV treatment with psoralens). Treatment begins with a minimal erythema dose-the lowest dose to achieve minimal erythema of the skin before burning. When that is determined, exposure is increased as needed—depending on the response. If this is impractical or too time-consuming for the patient, an alternative recommendation would be increased exposure to natural sunlight or even use of a tanning booth. However, patients must then be cautioned about the increased risk of skin cancer.

Refractory/severe psoriasis

Patient B is a 35-year-old man with a longstanding history of psoriasis affecting his scalp and nails. Over the past 10 years, psoriatic le-

TABLE Biologic agents for treating psoriasis²²

Agent (brand name; mechanism of action)	Maintenance dosing schedule	Baseline lab work (maintenance)	Adverse effects	Pertinent patient history	Approved uses
Adalimumab (Humira; TNF-α inhibitor)	SQ q2 wk	HBV, HCV, PPD/ interferon gamma release assay, CBC, CMP; also consider HIV test in high- risk individuals (PPD/interferon gamma release assay annually)	Risk of URI, injection site reaction, reactivation of TB, psoriatic reaction	Ask about prior/current malignancy, CHF, immunosuppression, hypersensitivity reactions, current/frequent infections, family or personal history of demyelinating disease (MS)	Rheumatoid arthritis, psoriatic arthritis, ankylosing spondylitis, juvenile idiopathic arthritis, Crohn disease, ulcerative colitis, plaque psoriasis, hidradenitis suppurativa, uveitis
Golimumab (Simponi; TNF-α inhibitor)	SQ q8 wk				Psoriatic arthritis, rheumatoid arthritis, ankylosing spondylitis, ulcerative colitis
Etanercept (Enbrel; TNF- α inhibitor)	SQ q1 wk				Rheumatoid arthritis, plaque psoriasis, psoriatic arthritis, ankylosing spondylitis, juvenile idiopathic arthritis for children ≥ 4 y
Ustekinumab (Stelara; anti-IL-12 and IL-23)	SQ q90 d				Plaque psoriasis, psoriatic arthritis, Crohn disease for children ≥ 12 y
Secukinumab (Cosentyx; anti-IL-17 and IL-22)	SQ q4 wk		Same as above, plus risk of IBD		Plaque psoriasis, psoriatic arthritis, ankylosing spondylitis
Ixekizumab (Taltz; anti-IL-17)	SQ q2 wk		pius risk Oi IBD	Same as above, plus history of IBD	Psoriatic arthritis, plaque psoriasis
Guselkumab (Tremfya; anti-IL-23)	SQ q8 wk	PPD/interferon gamma release assay	Risk of infection, injection site reaction, GI effects		Plaque psoriasis

CBC, complete blood count; CHF, congestive heart failure; CMP, comprehensive metabolic panel; GI, gastrointestinal; HBV, hepatitis B virus; HCV, hepatitis C virus; IBD, inflammatory bowel disease; IL, interleukin; MS, multiple sclerosis; PPD, purified protein derivative; SQ, subcutaneous injection; TB, tuberculosis; TNF, tumor necrosis factor; URI, upper respiratory infection.

sions have also appeared and grown across his lower back, gluteal fold, legs, abdomen, and arms. He is now being evaluated by a rheumatologist for worsening symmetric joint pain that includes his lower back.

■ Methotrexate has been used to treat psoriasis and psoriatic arthritis since the 1950s. Methotrexate is a competitive inhibitor of dihydrofolate reductase and is typically given as an oral medication dosed once weekly with folic acid supplementation on the other 6 days. ¹⁷ The most common adverse effects encountered with methotrexate are gastrointestinal upset and oral ulcers; how-

ever, routine monitoring for myelosuppression and hepatotoxicity is required.

■ Biologic therapy. When conventional therapies fail, immune-targeted treatment with "biologics" may be initiated. As knowledge of signaling pathways and the immunopathogenesis of psoriasis has increased, so has the number of biologic agents, which are generally well tolerated and effective in managing plaque psoriasis and psoriatic arthritis. Although their use, which requires monitoring, is handled primarily by specialists, familiarizing yourself with available agents can be helpful (TABLE).²²

CONTINUED



Individualize

according to

the severity of

of the lesions,

involvement

of joints, and

comorbidities.

disease, location

therapy

Nutritional modification and supplementation in treating skin disease still requires further investigation. Fish oil has shown benefit for cutaneous psoriasis in randomized controlled trials.^{7,8} Oral vitamin D supplementation requires further study, whereas selenium and B₁₂ supplementation have not conferred consistent benefit.⁷ Given that several studies have demonstrated a relationship between body mass index and psoriatic disease severity, weight loss may be helpful in the management of psoriasis as well as psoriatic arthritis.⁸

Other systemic agents—for individuals who cannot tolerate the biologic agents—include acitretin, azathioprine, mycophenolate mofetil, and cyclosporine. 15,17

Paradoxical psoriatic reactions

When a psoriatic condition develops during biologic drug therapy, it is known as a paradoxical psoriatic reaction. The onset of de novo psoriasis has been documented during TNF- α inhibitor therapy for individuals with underlying rheumatoid arthritis.²³ Skin biopsy reveals the same findings as common plaque psoriasis.

Using immunosuppressive Tx? Screen for tuberculosis

Testing to exclude a diagnosis of latent or undiagnosed tuberculosis must be performed prior to initiating immunosuppressive therapy with methotrexate or a biologic agent. Tuberculin skin testing, QuantiFERON-TB gold test, and the T-SPOT.TB test are accepted screening modalities. Discordance between tuberculin skin tests and the interferon gamma release assays in latent TB highlights the need for further study using the available QuantiFERON-TB gold test and the T-SPOT.TB test.²⁴

CORRESPONDENCE

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