# Management of Psoriasis With Topicals: Applying the 2020 AAD-NPF Guidelines of Care to Clinical Practice

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#### PRACTICE **POINTS**

- Topical medications collectively represent the most common form of psoriasis treatment. Depending on disease severity and distribution, topical agents can be used as monotherapy or adjunct therapy, offering the benefit of localized treatment without systemic side effects.
- Dermatologists should base the selection of an appropriate topical medication on factors including adverse effects, potency, vehicle, and anatomic localization of disease.

Topical medications have high utility in the treatment of psoriasis because of their localized effect and ability to be used as both monotherapy and adjunctive therapy. The American Academy of Dermatology (AAD) and the National Psoriasis Foundation (NPF) published guidelines in 2020 regarding the management of psoriasis with topical therapies. These guidelines are a framework that assist clinicians treating psoriasis patients with topical agents including steroids, calcineurin inhibitors (CNIs), vitamin D analogues, retinoids (tazarotene), emollients, keratolytics (salicylic acid), anthracenes (anthralin), and keratoplastics (coal tar). This review presents these evidence-based recommendations in a form that dermatologists can readily apply to their clinical practice. The selection of an appropriate topical therapy, effective combination therapies, duration of use, and adverse events are addressed.

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P soriasis is a chronic inflammatory skin disease characterized by erythematous scaly plaques that can invoke substantial pain, pruritus, and qualityof-life disturbance in patients. Topical therapies are the most commonly used medications for treating psoriasis, with one study (N=128,308) showing that more than 85% of patients with psoriasis were managed solely with topical medications.<sup>1</sup> For patients with mild to moderate psoriasis, topical agents alone may be able to control disease completely. For those with more severe disease, topical agents are used adjunctively with systemic or biologic agents to optimize disease control in localized areas.

The American Academy of Dermatology (AAD) and National Psoriasis Foundation (NPF) published guidelines in 2020 for managing psoriasis with topical agents in adults.<sup>2</sup> This review presents the most up-to-date clinical recommendations for topical agent use in adult

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patients with psoriasis and elaborates on each drug's pharmacologic and safety profile. Specifically, evidencebased treatment recommendations for topical steroids, calcineurin inhibitors (CNIs), vitamin D analogues, retinoids (tazarotene), emollients, keratolytics (salicylic acid), anthracenes (anthralin), and keratoplastics (coal tar) will be addressed (Table 1). Recommendations for combination therapy with other treatment modalities including UVB light therapy, biologics, and systemic nonbiologic agents also will be discussed.

#### Selecting a Topical Agent Based on Disease Localization

When treating patients with psoriasis with topical therapies, clinicians should take into consideration drug potency, as it determines how effective a treatment will be in penetrating the skin barrier. Plaque characteristics, such as distribution (localized vs widespread), anatomical localization (flexural, scalp, palms/soles/nails), size (large vs small), and thickness (thick vs thin), not only influence treatment effectiveness but also the incidence of drug-related adverse events. Furthermore, preferred topical therapies are tailored to each patient based on disease characteristics and activity. Coal tar and anthralin have been used less frequently than other topical therapies for psoriasis because of their undesirable side-effect profiles (Table 1).<sup>3</sup>

*Face and Intertriginous Regions*—The face and intertriginous areas are sensitive because skin tends to be thin in these regions. Emollients are recommended for disease in these locations given their safety and flexibility in use for most areas. Conversely, anthralin should be avoided on the face, intertriginous areas, and even highly visible locations because of the potential for skin staining. Low-potency corticosteroids also have utility in psoriasis distributed on the face and intertriginous regions. Additionally, application of steroids around the eyes should be cautioned because topical steroids can induce ocular complications such as glaucoma and cataracts in rare circumstances.<sup>4</sup>

Off-label use of CNIs for psoriasis on the face and intertriginous areas also is effective. Currently, there is a level B recommendation for off-label use of 0.1% tacrolimus for up to 8 weeks for inverse psoriasis or psoriasis on the face. Off-label use of pimecrolimus for 4 to 8 weeks also can be considered for inverse psoriasis. Combination therapy consisting of hydrocortisone with calcipotriol ointment is another effective regimen.<sup>5</sup> One study also suggested that use of crisaborole for 4 to 8 weeks in intertriginous psoriasis can be effective and well tolerated.<sup>6</sup>

*Scalp*—The vehicle of medication administration is especially important in hair-bearing areas such as the scalp, as these areas are challenging for medication application and patient adherence. Thus, patient preferences for the vehicle must be considered. Several studies have been conducted to assess preference for various

vehicles in scalp psoriasis. A foam or solution may be preferable to ointments, gels, or creams.<sup>7</sup> Gels may be preferred over ointments.<sup>8</sup> There is a level A recommendation supporting the use of class 1 to 7 topical steroids for a minimum of 4 weeks as initial and maintenance treatment of scalp psoriasis. The highest level of evidence (level A) also supports the use of calcipotriol foam or combination therapy of calcipotriol–betamethasone dipropionate gel for 4 to 12 weeks as treatment of mild to moderate scalp psoriasis.

*Nails*—Several options for topical medications have been recommended for the treatment of nail psoriasis. Currently, there is a level B recommendation for the use of tazarotene for the treatment of nail psoriasis. Another effective regimen is combination therapy with vitamin D analogues and betamethasone dipropionate.<sup>9</sup> Topical steroid use for nail psoriasis should be limited to 12 weeks because of the risk for bone atrophy with chronic steroid use.

*Palmoplantar*—The palms and soles have a thicker epidermal layer than other areas of the body. As a result, class 1 corticosteroids can be used for palmoplantar psoriasis for more than 4 weeks with vigilant monitoring for adverse effects such as skin atrophy, tachyphylaxis, or tinea infection. Tazarotene also has been shown to be helpful in treating palmoplantar psoriasis.

*Resistant Disease*—Intralesional steroids are beneficial treatment options for recalcitrant psoriasis in glabrous areas, as well as for palmoplantar, nail, and scalp psoriasis. Up to 10 mg/mL of triamcinolone acetonide used every 3 to 4 weeks is an effective regimen.<sup>10</sup>

Pregnancy/Breastfeeding—Women of childbearing potential have additional safety precautions that should be considered during medication selection. Emollients have been shown to be safe during pregnancy and lactation. Currently, there is little known about CNI use during pregnancy. During lactation, CNIs can be used by breastfeeding mothers in most areas, excluding the breasts. Evaluation of the safety of anthralin and vitamin D analogues during pregnancy and lactation have not been studied. For these agents, dermatologists need to use their clinical judgment to weigh the risks and benefits of medication, particularly in patients requiring occlusion, higher medication doses, or treatment over a large surface area. Salicylic acid should be used with caution in pregnant and breastfeeding mothers because it is a pregnancy category C drug. Lower-potency corticosteroids may be used with caution during pregnancy and breastfeeding. More potent corticosteroids and coal tar, however, should be avoided. Similarly, tazarotene use is contraindicated in pregnancy. According to the US Food and Drug Administration labels for all forms of topical tazarotene, a pregnancy test must be obtained 2 weeks prior to tazarotene treatment initiation in women of childbearing potential because of the risk for serious fetal malformations and toxicity.

| Therapy                   | Recommended duration   | Frequency                | Recommendations (level of strength <sup>a</sup> )   | Common adverse events  |
|---------------------------|--|--------------------------|---|--|
| Steroids                  | 2–4 wk for mild<br>to severe plaque<br>psoriasis; class 1 can<br>be used on palms<br>and soles for >4 wk<br>with monitoring of<br>side effects | Twice daily <sup>b</sup> | Use of class 1–5 recommended for<br>treatment of plaque psoriasis with<br>the exception of intertriginous areas<br>(level A); use of class 1–7 for $\geq$ 4 wk is<br>recommended for initial and maintenance<br>treatment of scalp psoriasis (level A);<br>mid-potency or high-potency steroid in<br>combination with tazarotene for 8–16 wk<br>for mild to moderate psoriasis is more<br>effective than tazarotene monotherapy<br>(level A); topical steroid in combination<br>with tazarotene is recommended to<br>decrease the length of treatment and<br>increase disease remission time (level A);<br>class 1 topical steroid in combination with<br>etanercept for 12 wk is recommended<br>for moderate to severe psoriasis (level A);<br>use of salicylic acid in combination with<br>topical corticosteroids to treat moderate<br>to severe psoriasis (level B) | Skin atrophy,<br>telangiectasia,<br>folliculitis, purpura,<br>and contact<br>dermatitis;<br>exacerbation of<br>acne, rosacea,<br>perioral dermatitis,<br>and tinea<br>infection; rebound<br>phenomenon if not<br>tapered |
| Calcineurin<br>inhibitors | Can be used<br>for >4 wk   | Twice daily              | Off-label use of 0.1% tacrolimus for inverse<br>psoriasis or psoriasis on the face for up to<br>8 wk (level B); off-label use of pimecrolimus<br>for inverse psoriasis for 4–8 wk (level B);<br>off-label use of tacrolimus in combination<br>with 6% salicylic acid for 12 wk for plaque<br>psoriasis (level B)  | Burning, pruritus,<br>flushing with<br>ingestion of alcohol  |
| Vitamin D<br>analogues    | Safe for long-term use<br>(recommended for up<br>to 52 wk); use for<br>4–8 wk is effective<br>for mild to moderate<br>psoriasis                | Twice daily              | Up to 52-wk usage of topical vitamin D<br>analogues is recommended for mild to<br>moderate psoriasis (level A); calcipotriol<br>foam and calcipotriol–betamethasone<br>dipropionate gel for 4–12 wk for<br>scalp psoriasis (level A); calcipotriol in<br>combination with class 2 and 3 topical<br>steroids for up to 52 wk is recommended<br>for treatment of psoriasis (level A);<br>combination treatment with calcipotriol<br>and corticosteroids is recommended<br>(level A); topical calcipotriol in combination<br>with methotrexate for treatment of<br>moderate to severe psoriasis (level A);<br>calcipotriol + acitretin for moderate to<br>severe psoriasis (level A); calcipotriol–<br>betamethasone dipropionate ointment<br>+ low-dose cyclosporine for moderate to<br>severe psoriasis (level B)  | Burning, pruritus,<br>erythema, edema,<br>peeling, dryness   |
| Tazarotene                | Use for 8–12 wk for<br>mild to moderate<br>psoriasis   | Once daily               | Treatment for mild to moderate psoriasis<br>(level B); treatment of nail psoriasis<br>(level B); combination of tazarotene and<br>NB-UVB is effective and reduces amount<br>of NB-UVB needed (level B)  | Erythema, burning, pruritus  |
|                           |  |                          |   | CONTINUED ON NEXT PAGE   |

## TABLE 1. Summary of Topical Therapies for Adults With Psoriasis

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TABLE 1. (continued)

| Therapy        | Recommended duration                                 | Frequency   | Recommendations (level of strength®)   | Common adverse<br>events   |
|----------------|--|---|--|--|
| Emollients     | Any  | No<br>consistent<br>frequency<br>in the<br>literature | Emollient + topical steroids for 4–8 wk<br>(level B)   | Contact dermatitis   |
| Salicylic acid | Use for 8–16 wk for<br>mild to moderate<br>psoriasis | No<br>consistent<br>frequency<br>in the<br>literature | Treatment for mild to moderate psoriasis<br>(level B)  | Salicylate toxicity  |
| Anthralin      | Use for 8–12 wk for<br>mild to moderate<br>psoriasis | No<br>consistent<br>frequency<br>in the<br>literature | Treatment for mild to moderate psoriasis;<br>to minimize side effects, application should<br>be limited to 2 h/d (level B)   | Erythema, burning,<br>staining of skin   |
| Coal tar       | N/A  | No<br>consistent<br>frequency<br>in the<br>literature | Coal tar is recommended for treatment<br>of mild to moderate psoriasis (level A);<br>Goeckerman therapy can be recommended<br>for treatment of psoriasis (level B) | Irritation, folliculitis,<br>contact dermatitis,<br>phototoxicity, tar<br>pigmentation, odor |

Abbreviations: FDA, US Food and Drug Administration; N/A, not applicable; NB-UVB, narrowband UVB.

<sup>a</sup>Strength of Recommendation Taxonomy (SORT): A, recommendation based on consistent and good-quality, patient-oriented evidence; B, recommendation based on inconsistent or limited-quality, patient-oriented evidence; C, recommendation based on consensus, opinion, case studies, or disease-oriented evidence.

<sup>b</sup>Triamcinolone acetonide 0.025% cream or ointment is FDA approved for use 2–4 times daily but is most commonly used 2 times daily. Triamcinolone acetonide 0.1% and 0.5% cream or ointment are FDA approved for use 2–3 times daily but are most commonly used 2 times daily.

#### Recommendations, Risks, and Benefits of Topical Therapy for the Management of Psoriasis

*Topical Corticosteroids*—Topical corticosteroids (TCs) are widely used for inflammatory skin conditions and are available in a variety of strengths (Table 2). They are thought to exert their action by regulating the gene transcription of proinflammatory mediators. For psoriasis, steroids are recommended for 2 to 4 weeks, depending on disease severity. Although potent and superpotent steroids are more effective than mild- to moderate-strength TCs, use of lower-potency TCs may be warranted depending on disease distribution and localization.<sup>11</sup> For treatment of psoriasis with no involvement of the intertriginous areas, use of class 1 to 5 TCs for up to 4 weeks is recommended.

For moderate to severe psoriasis with 20% or less body surface area (BSA) affected, combination therapy consisting of mometasone and salicylic acid has been shown to be more effective than mometasone alone.<sup>12,13</sup> There currently is a level A recommendation for the use of combination therapy with class 1 TCs and etanercept for 12 weeks in patients with moderate to severe psoriasis who require both systemic and topical therapies for disease control. Similarly, combination therapy with infliximab and high-potency TCs has a level B recommendation to enhance efficacy for the treatment of moderate to severe psoriasis.<sup>14</sup> High-quality studies on the use of TCs with anti–IL-12/IL-23, anti–IL-23, and anti–IL-17 currently are unavailable, but the combination is not expected to be unsafe.<sup>14,15</sup> Combination therapy of betamethasone dipropionate ointment and low-dose cyclosporine is an alternative regimen with a level B recommendation.

The most common adverse effects with use of TCs are skin thinning and atrophy, telangiectasia, and striae (Table 1). With clinical improvement of disease, it is recommended that clinicians taper TCs to prevent rebound effect. To decrease TC-related adverse effects, clinicians should use combination therapy with steroid-sparing agents for disease maintenance, transition to lower-potency corticosteroids, or use intermittent steroid therapy. Systemic effects of TC use include hypothalamic-pituitary-adrenal

axis suppression, Cushing syndrome, and osteonecrosis of the femoral head.<sup>16-18</sup> These systemic effects with TC use are rare unless treatment is for disease involving greater than 20% BSA or occlusion for more than 4 weeks.

*Calcineurin Inhibitors*—Calcineurin inhibitors inhibit calcineurin phosphorylation and T-cell activation, subsequently decreasing the expression of proinflammatory cytokines. Currently, they are not approved by the US Food and Drug Administration to treat psoriasis but have demonstrated efficacy in randomized control trials (RCTs) for facial and intertriginous psoriasis. In RCTs, 71% of patients using pimecrolimus cream 0.1% twice daily for 8 weeks achieved an investigator global assessment score of clear (0) or almost clear (1) compared with 21% of placebo-treated patients (N=57).<sup>19</sup> Other trials have shown that 65% of patients receiving tacrolimus ointment 0.1% for 8 weeks achieved an investigator global assessment score of 0 or 1 compared with 31% of placebo-treated patients (N=167).<sup>20</sup> Because of their efficacy in RCTs, CNIs commonly are used off label to treat psoriasis.

The most common adverse effects with CNI use are burning, pruritus, and flushing with alcohol ingestion (Table 1). Additionally, CNIs have a black box warning that use may increase the risk for malignancy, but this risk has not been demonstrated with topical use in humans.<sup>21</sup>

*Vitamin D Analogues*—The class of vitamin D analogues—calcipotriol/calcipotriene and calcitriol—frequently are used to treat psoriasis.Vitamin D analogues exert their beneficial effects by inhibiting keratinocyte proliferation and enhancing keratinocyte differentiation. They also are ideal for long-term use (up to 52 weeks) in mild to moderate psoriasis and can be used in combination with class 2 and 3 TCs. There is a level A recommendation that supports the use of combination therapy with calcipotriol and TCs for the treatment of mild to moderate psoriasis.

| Class  | Corticosteroid  |  |  |
|--|---|--|--|
| 1 (superpotent)  | Augmented betamethasone dipropionate 0.05% gel or ointment; augmented diflorasone diacetate 0.05% ointment; clobetasol propionate 0.05% in any vehicle; desoximetasone 0.25% spray; fluocinonide 0.1% cream; flurandrenolide 4 µg/cm <sup>2</sup> tape; halobetasol propionate 0.05% cream or ointment  |  |  |
| 2 (high potency)   | Amcinonide 0.1% ointment; augmented betamethasone dipropionate 0.05% cream or lotion; augmented diflorasone diacetate 0.05% cream; betamethasone dipropionate 0.05% ointment; desoximetasone 0.05% gel; desoximetasone 0.25% cream or ointment; diflorasone diacetate 0.05% ointment; fluocinonide 0.05% cream, gel, or ointment; halcinonide 0.1% cream or ointment; mometasone furoate 0.1% ointment; triamcinolone acetonide 0.5% ointment   |  |  |
| 3 (high potency)   | Amcinonide 0.1% cream or lotion; betamethasone dipropionate 0.05% cream; betamethasor valerate 0.1% ointment; diflorasone diacetate 0.05% cream; fluticasone propionate 0.005% ointment; triamcinolone acetonide 0.1% ointment; triamcinolone acetonide 0.5% cream  |  |  |
| 4 (medium potency)   | Betamethasone valerate 0.12% foam; desoximetasone 0.05% cream; fluocinolone acetonic<br>0.025% ointment; fluticasone propionate 0.05% cream; flurandrenolide 0.05% ointment;<br>hydrocortisone valerate 0.2% ointment; mometasone furoate 0.1% cream or lotion; triamcir<br>acetonide 0.1% cream; triamcinolone acetonide 0.2% spray  |  |  |
| 5 (medium potency)   | Betamethasone dipropionate 0.05% lotion; betamethasone valerate 0.1% cream or lotion; clocortolone pivalate 0.1% cream; fluocinolone acetonide 0.025% cream; fluocinolone acetonide 0.01% oil or shampoo; fluticasone propionate 0.05% cream or lotion; flurandrenolide 0.05% cream or lotion; hydrocortisone butyrate 0.1% cream, ointment, lotion, or solution; hydrocortisone probutate cream 0.1%; hydrocortisone valerate cream 0.2%; prednicarbate 0.1% cream or ointment; triamcinolone acetonide 0.025% ointment; triamcinolone acetonide 0.1% lotion |  |  |
| 6 (low potency) Alclometasone dipropionate 0.05% cream or ointment; betamethasone valerate 0 desonide 0.05% in any vehicle; fluocinolone acetonide 0.01% cream or solution; t acetonide 0.025% cream or lotion |   |  |  |
| 7 (low potency)  | Dexamethasone sodium phosphate 0.1% cream; hydrocortisone 0.5%–2.5% cream, ointment, gel, lotion, or solution; methylprednisolone acetate 0.25% cream   |  |  |

### **TABLE 2.** Topical Corticosteroids by Class

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For severe psoriasis, many studies have investigated the efficacy of combination therapy with vitamin D analogues and systemic treatments. Combination therapy with calcipotriol and methotrexate or calcipotriol and acitretin are effective treatment regimens with level A recommendations. Calcipotriolbetamethasone dipropionate ointment in combination with low-dose cyclosporine is an alternative option with a level B recommendation. Because vitamin D analogues are inactivated by UVA and UVB radiation, clinicians should advise their patients to use vitamin D analogues after receiving UVB phototherapy.<sup>22</sup>

Common adverse effects of vitamin D analogues include burning, pruritus, erythema, and dryness (Table 1). Hypercalcemia and parathyroid hormone suppression are extremely rare unless treatment occurs over a large surface area (>30% BSA) or the patient has concurrent renal disease or impairments in calcium metabolism.

*Tazarotene*—Tazarotene is a topical retinoid that acts by decreasing keratinocyte proliferation, facilitating keratinocyte differentiation, and inhibiting inflammation. Patients with mild to moderate psoriasis are recommended to receive tazarotene treatment for 8 to 12 weeks. In several RCTs, tazarotene gel 0.1% and tazarotene cream 0.1% and 0.05% achieved treatment success in treating plaque psoriasis.<sup>23,24</sup>

For increased efficacy, clinicians can recommend combination therapy with tazarotene and a TC. Combination therapy with tazarotene and a mid- or high-potency TC for 8 to 16 weeks has been shown to be more effective than treatment with tazarotene alone.<sup>25</sup> Thus, there is a level A recommendation for use of this combination to treat mild to moderate psoriasis. Agents used in combination therapy work synergistically to decrease the length of treatment and increase the duration of remission. The frequency of adverse effects, such as irritation from tazarotene and skin atrophy from TCs, also are reduced.<sup>26</sup> Combination therapy with tazarotene and narrowband UVB (NB-UVB) is another effective option that requires less UV radiation than NB-UVB alone because of the synergistic effects of both treatment modalities.27 Clinicians should counsel patients on the adverse effects of tazarotene, which include local irritation, burning, pruritus, and erythema (Table 1).

*Emollients*—Emollients are nonmedicated moisturizers that decrease the amount of transepidermal water loss. There is a level B recommendation for use of emollients and TCs in combination for 4 to 8 weeks to treat psoriasis. In fact, combination therapy with mometasone and emollients has demonstrated greater improvement in symptoms of palmoplantar psoriasis (ie, erythema, desquamation, infiltration, BSA involvement) than mometasone alone.<sup>28</sup> Emollients are safe options that can be used on all areas of the body and during pregnancy and lactation. Although adverse effects of emollients are rare, clinicians should counsel patients on the risk for contact dermatitis if specific allergies to ingredients/fragrances exist (Table 1).

*Salicylic Acid*—Salicylic acid is a topical keratolytic that can be used to treat psoriatic plaques. Use of salicylic acid for 8 to 16 weeks has been shown to be effective for mild to moderate psoriasis. Combination therapy of salicylic acid and TCs in patients with 20% or less BSA affected is a safe and effective option with a level B recommendation. Combination therapy with salicylic acid and calcipotriene, however, should be avoided because calcipotriene is inactivated by salicylic acid. It also is recommended that salicylic acid application follow phototherapy when both treatment modalities are used in combination.<sup>29,30</sup> Clinicians should be cautious about using salicylic acid in patients with renal or hepatic disease because of the increased risk for salicylate toxicity (Table 1).

Anthralin—Anthralin is a synthetic hydrocarbon derivative that has been shown to reduce inflammation and normalize keratinocyte proliferation through an unknown mechanism. It is recommended that patients with mild to moderate psoriasis receive anthralin treatment for 8 to 12 weeks, with a maximum application time of 2 hours per day. Combination therapy of excimer laser and anthralin has been shown to be more effective in treating psoriasis than anthralin alone.<sup>31</sup> Therefore, clinicians have the option of including excimer laser therapy for additional disease control. Anthralin should be avoided on the face, flexural regions, and highly visible areas because of potential skin staining (Table 1). Other adverse effects include application-site burning and erythema.

*Coal Tar*—Coal tar is a heterogenous mixture of aromatic hydrocarbons that is an effective treatment of psoriasis because of its inherent anti-inflammatory and keratoplastic properties. There is high-quality evidence supporting a level A recommendation for coal tar use in mild to moderate psoriasis. Combination therapy with NB-UVB and coal tar (also known as Goeckerman therapy) is a recommended treatment option with a quicker onset of action and improved outcomes compared with NB-UVB therapy alone.<sup>32,33</sup> Adverse events of coal tar include application-site irritation, folliculitis, contact dermatitis, phototoxicity, and skin pigmentation (Table 1).

#### Conclusion

Topical medications are versatile treatment options that can be utilized as monotherapy or adjunct therapy for mild to severe psoriasis. Benefits of topical agents include minimal required monitoring, few contraindications, and direct localized effect on plaques. Therefore, side effects with topical agent use rarely are systemic. Medication interactions are less of a concern with topical therapies; thus, they have better safety profiles compared with systemic therapies. This clinical review

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summarizes the recently published evidence-based guidelines from the AAD and NPF on the use of topical agents in psoriasis and may be a useful guiding framework for clinicians in their everyday practice.

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