PSORIASIS (JAISHIN WU, SECTION EDITOR)



A Review and Update of Phototherapy Treatment Options for Psoriasis

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Abstract

Purpose of Review Psoriasis is a chronic inflammatory skin condition affecting greater than 7 million adults in the USA. There are a multitude of treatment options available for psoriasis management. This review provides an overview of the current use of phototherapy in the treatment of psoriasis, and an update on recent phototherapy advances, either as a single modality or in conjunction with other agents.

Recent Findings Advances in pharmacotherapy and an increased understanding of the pathogenesis of psoriasis have led to the development of new systemic treatments for psoriasis. A small number of recent studies have evaluated the use of phototherapy in comparison to or in combination with emerging and traditional treatment modalities.

Summary Psoriasis is a chronic skin condition with a high disease burden among patients. More than half of patients are dissatisfied with their therapy, and many are not receiving adequate levels of treatment. Phototherapy remains an effective modality for the treatment of psoriasis with minimal risk. While some studies have evaluated the use of phototherapy with systemic options, more studies are needed to make definitive conclusions.

Keywords Psoriasis · Phototherapy · Ultraviolet A (UVA) · Ultraviolet B (UVB) · Narrowband ultraviolet B (NB-UVB) · Broadband ultraviolet B (BB-UVB) · Psoralen ultraviolet A (PUVA) · Light-emitting diode (LED) · Excimer laser

Introduction

Psoriasis is a T cell-mediated disease that increases morbidity and mortality, reduces quality of life, and has a profound impact on patients' psychosocial well-being [1]. Psoriasis is categorized, based on the percentage of body surface area (BSA) involvement, as either mild (< 3% BSA), moderate (3-10%BSA), or severe (> 10% BSA). The impact upon the individual is also a consideration when evaluating psoriasis [2]. Treatment planning is developed based on these severity classifications and the impact of psoriasis on the individual (Fig. 1).

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Topical agents and localized phototherapy are ideal for mild psoriasis, while moderate-to-severe psoriasis is treated with topical therapy plus phototherapy and/or systemic therapy [3]. Topical therapy options include corticosteroids, vitamin D analogues, retinoids, anthralin, coal tar, and salicylic acid [4–7]. Systemic therapy options include non-biologics and biologics [8•].

The use of phototherapy for the treatment of skin conditions dates back thousands of years. Its utility in the treatment of psoriasis was established with the advent of broadband ultraviolet B (BB-UVB), narrowband ultraviolet B (NB-UVB), psoralen plus ultraviolet A (PUVA), and laser treatment modalities [9]. Phototherapy is considered a first-line therapy in the treatment of psoriasis due to its efficacy and limited contraindications [10]. We aim to provide an overview of the current use of phototherapy in the treatment of psoriasis and highlight recent updates in the field.

Methods

A PubMed search of clinical trials from May 1995 to November 2017, using the terms psoriasis, phototherapy, Fig. 1 Approach to psoriasis treatment. Options for psoriasis treatment based on disease severity. UVB ultraviolet B, PUVA psoralen plus ultraviolet



UVB, PUVA, UVA, excimer laser, and light-emitting diodes (LED), was conducted. This search generated articles pertaining to the use of phototherapy versus, or in combination with, topical, systemic, or psychological treatments (Table 1). Articles published within the last 5 years reporting patient-centered outcomes were selected for inclusion in this review. A separate PubMed search using the same terms was conducted without the use of a 5-year filter in order to generate additional information.

Phototherapy Overview

Mechanism of Action

Psoriasis is a T cell-mediated disease driven by T helper 1 (Th1) and T helper 17 (Th17) pathways. Keratinocyte hyperproliferation and inflammation occurs via a milieu of cytokines including tumor necrosis factor (TNF)- α , interleukin (IL)-17, and IL-23 [12••]. There are several mechanisms by which phototherapy may impact the pathogenesis of psoriasis

 Table 1
 Combination treatments.

 UVB plus additional therapeutic modalities
 Plus additional therapeutic modalities

Combination therapy options	Comments			
Tarazotene	Effective; improved efficacy over UVB alone			
Anthralin	Effective; cumbersome application			
Coal Tar	Effective; cumbersome and odor not always tolerated			
Vitamin D analogues	Conflicting data; must apply 2 h before or after phototherapy to avoid product degradation			
Topical steroids	Conflicting data; unclear at present if combination with UVB is beneficial			
Oral retinoids	Effective; reduced UVB response time and cumulative dose			
Methotrexate	May be beneficial in select patients			
Cyclosporine	Not recommended			
Alefacept	May be beneficial in select patients			
Etanercept	May be beneficial in select patients			
Additional biologics	Likely effective; limited data at present			
PUVA	Combination w/ NB-UVB may lead to faster clearing; potential for increased carcinogenicity limits its use			
Excimer laser	Combined use with UVB can be beneficial in increasing response time in areas with recalcitrant plaques			

Data is adapted from Lapolla et al. and Menter et al. [10, 11•]

UVB ultraviolet B, PUVA psoralen + ultraviolet A, NB-UVB narrowband ultraviolet B

[13]. Phototherapy shifts cytokine production in the direction of the counter-regulatory T helper 2 (Th2) immune response both locally and systemically [14, 15]. Keratinocyte apoptosis and upregulation of p53, a tumor suppressor gene, are induced by ultraviolet radiation [16, 17]. Phototherapy induces the migration of histiocytes out of the epidermis [18]. The cumulative effects of these mechanisms improve psoriasis [19].

Indications, Contraindications, and Adverse Reactions (See Table 2)

Phototherapy, either as a targeted or whole body treatment, improves mild or moderate-to-severe psoriasis, with NB-UVB constituting the most widely administered form [10, 20]. As with any treatment option, use should be tailored to the individual characteristics of the patient.

There are relatively few absolute contraindications to commencing UVB phototherapy or PUVA. Diagnoses of photosensitive dermatoses such as systemic lupus erythematous or xeroderma pigementosum are contraindications to phototherapy use [21]. Relative contraindications to phototherapy include personal history of melanoma or non-melanoma skin cancer, family history of melanoma, Fitzpatrick skin type 1/2, immune suppression due to solid organ transplant, emotional or physical limitation to treatment, or previous arsenic or ionizing radiation treatment. Pregnancy, lactation, liver disease, or a history of cyclosporine or methotrexate use are relative contraindications to the use of PUVA [11•, 21].

Although phototherapy is generally well tolerated, there are some short-term and long-term factors that may influence adherence and the phototherapy modality employed. Short-term complications include erythema, blistering, and itching. Longterm complications include photoaging and increased risk of developing non-melanoma skin cancer in those with higher cumulative doses of oral PUVA [11•, 22, 23]. Studies have not confirmed an increased risk of squamous or basal cell carcinoma with the use of UVB therapy alone [24].

Special Considerations

Pediatric and pregnant patients represent population subsets that often generate extra scrutiny prior to the administration of phototherapy. Pregnancy is not considered a contraindication to UVB phototherapy [25]. UVB phototherapy should be considered a primary treatment for pregnant patients with plaque or guttate psoriasis [26]. Phototherapy is a safe and effective second-line treatment option for pediatric patients who are unresponsive to topical therapy [27].

Table 2 Overview of phototherapy treatment. Phototherapy treatment considerations for NB-UVB, PUVA, and excimer laser

Туре	Indication	Efficacy	Absolute contraindications	Relative contraindications	Adverse effects
NB-UVB	 First line in moderate-severe plaque or guttate psoriasis Second line in mild psoriasis 	- 80% of patients clear with 3 times per week treatment	- Photosensitive dermatoses (i.e., SLE, xeroderma pigmentosa)	 Atypical nevi Non-melanoma skin cancers Personal/family history of melanoma Fitzpatrick skin types 	Acute: erythema, blistering, pruritus, hyperpigmentation, HSV recurrence Chronic: photoaging
				 1/2 Immune suppression due to organ transplant Physical or emotional inability to tolerate phototherapy Previous arsenic or ionizing radiation 	
PUVA	 Palmoplantar psoriasis Moderate-to-severe plaque psoriasis 	 Most efficacious form of phototherapy 90% of patients clear within 30 treatments 	 Photosensitive dermatoses (i.e., SLE, xeroderma pigmentosa) Porphyria 	- Pregnant or nursing - History of treatment with cyclosporine or methotrexate	Acute: nausea, erythema, pruritus, hyperpigmentation, HSV recurrence, CNS disturbance Chronic: photodamage, dose-dependent risk of NMSC
Excimer laser	- Localized psoriasis	- Plaque clearance occurs within 12 treatments generally		- Photosensitive dermatoses (i.e., SLE, xeroderma pigmentosa)	Acute: erythema, blistering Chronic: hyperpigmentation

NB-UVB narrowband ultraviolet B, SLE systemic lupus erythematosus, HSV herpes simplex virus, PUVA psoralen + ultraviolet A, CNS central nervous system, NMSC non-melanotic skin cancer

Home Phototherapy

The time commitment and potential cost of office-based phototherapy can be prohibitive for many patients. The potential success of a phototherapy regimen may, in part, depend on the location of treatment [28]. Home UVB is as effective as office-based UVB and is associated with a greater degree of patient satisfaction [29]. In appropriate patients, home UVB may be an ideal option for the treatment of psoriasis [8•].

Clinical Use

Prior to initiating treatment with phototherapy, psoriasis disease severity should be assessed. Objective assessments include psoriasis area and severity index (PASI), BSA, and physician global assessment (PGA) [19]. Phototherapy protocols are typically based on the minimal erythema dose (MED) or the Fitzpatrick skin type [11•]. The MED is the lowest dose of UVB that generates trace erythema 24 h after commencing UVB therapy [30]. BB-UVB, NB-UVB, excimer laser, and PUVA protocols recommended by the American Academy of Dermatology are available online [31–34].

Broadband Ultraviolet B, Narrowband Ultraviolet B, and Excimer Laser

NB-UVB (311–313 nm) is a first-line treatment for patients with moderate-to-severe psoriasis and a second-line treatment for patients with mild disease which has been unresponsive to topical therapy. BB-UVB (290–320 nm) treatments begin with an initial 20- to 60-mJ/cm² dose based on the Fitzpatrick skin type, or a dose equivalent to 50% of MED, and are typically administered three to five times per week [11•].

NB-UVB has largely replaced the use of BB-UVB due to decreased time to the achievement of plaque clearance, and an increased duration of remission [35]. The side effects of BB-UVB and NB-UVB are similar. Since efficacy of NB-UVB is achieved at a lower total cumulative dose of UVB radiation, the long-term complications of UVB phototherapy may be reduced when using NB-UVB [36, 37]. NB-UVB treatments begin with an initial 130- to 400-mJ/cm² dose based on the Fitzpatrick skin type, or a dose equivalent to 50% of MED, and are typically administered three to five times per week [11•].

Excimer, or "excited dimer," laser therapy is a targeted phototherapy treatment which uses 308 nm UVB light, and has emerged over the last two decades as an effective option for localized disease unresponsive to topical steroids [38]. Excimer treatment, and other forms of localized UV treatment, has the benefit of targeting only involved skin, while sparing uninvolved areas, allowing for a reduction in the number of treatments and the long-term adverse effects of phototherapy [39, 40]. Localized UV is also useful in treating recalcitrant areas and locations which are difficult to reach with generalized phototherapy [41]. Excimer therapy is initiated based on skin type and plaque thickness, at a dose range of 400 to 900 mJ/cm²; treatment is typically administered two to three times per week [11•].

Psoralen and Ultraviolet A

PUVA is an established and efficacious form of phototherapy and is particularly useful in the treatment of palmoplantar psoriasis. However, concerns over increased long-term risk of carcinogenicity have led to reduction in its use at present [42]. UVA (340–400 nm) penetrates deeper into the skin than UVB and modifies processes deeper in the dermal layer by effecting blood vessels, endothelial cells, dermal fibroblasts, and dendritic cells in addition to reducing keratinocyte proliferation and promoting histiocyte migration out of the epidermis [43].

UVA therapy can be combined with oral or topical psoralen to produce a greater degree of immune suppression than observed with UVA alone [44]. Standard therapy in the USA is to administer oral 8-methoxypsoralen which intercalates between DNA base pairs. When subjected to UVA, 8methoxypsoralen inhibits DNA replication by forming crosslinks. The activation of psoralen by UVA also promotes membrane and mitochondrial damage by promoting the formation of reactive oxygen species [45]. 8-Methoxypsoralen is administered 1 to 2 h prior to UVA treatment at a dose of 0.4– 0.6 mg/kg. Treatment is typically administered two to three times per week [11•].

Current Trends and Recent Updates

NB-UVB

NB-UVB in Combination with Behavioral Therapy

There is some evidence to support the idea that psychological stress may reduce the rate of clearance in psoriasis patients undergoing treatment with phototherapy [46, 47]. A study evaluating response to NB-UVB in 40 subjects randomized to either NB-UVB or NB-UVB plus cognitive behavioral therapy (CBT) noted that the PASI75 rate at 8 weeks in the NB-UVB plus the CBT group was higher than the NB-UVB group alone (65 vs. 15%; p = 0.007) [48].

NB-UVB in Combination with Vitamin D Analogues

Vitamin D analogues, such as calcipotriene, when used concurrently with UVB phototherapy decrease the time to

symptomatic relief and lower the overall cumulative UVB dose that patients receive [49]. A 12-week, open-label, prospective right-left, intra-individual clinical trial assessed the efficacy and safety of topical tacalcitol in combination with NB-UVB versus NB-UVB alone in 30 subjects with psoriasis. Target lesions on one side of the subjects were treated with tacalcitol ointment; no treatment was applied to target lesions on the opposite side. NB-UVB was administered three times weekly. Tacalcitol plus NB-UVB use led to earlier plaque clearance and better maintenance of response than NB-UVB alone [50]. In comparing the efficacy of calcipotriol versus tacalcitol in combination with NB-UVB, 30 subjects with stable plaque psoriasis were enrolled in 12-week clinical trial. Target lesions on the subject's left side were treated with tacalcitol ointment once daily while target lesions on the subject's right side were treated with calcipotriol ointment twice daily. NB-UVB was administered three times weekly [51]. The calcipotriol group achieved lower mean target plaque scores throughout the study (p < 0.05). At 12 weeks, 90.0% of subjects in the calcipotriol group versus 76.67% of subjects in the tacalcitol group achieved or maintained target plaque clearance (p < 0.05). Adverse events (AEs) were mild, consisting mainly of pruritus and hyperpigmentation, and were similar among the two groups [51].

NB-UVB in Combination with Non-biologic Systemic Therapies

In a recent study, the efficacy of methotrexate (MTX) plus NB-UVB versus either MTX or NB-UVB alone was evaluated in 113 subjects with psoriasis. The MTX plus NB-UVB group achieved PASI90 at 6.11 ± 1.28 versus 20.87 ± 4.21 weeks in the MTX alone group, and 11.42 ± 2.36 in the NB-UVB alone group (p < 0.0001). The cumulative dose of NB-UVB was 12.13 ± 4.02 J/cm² in the MTX plus NB-UVB group versus 34.48 ± 13.13 J/cm² in the NB-UVB alone group (p < 0.0001). The cumulative dose of MTX was 116.04 ± 20.47 mg in the MTX + NB-UVB group versus 298.63 ± 60.26 in the MTX alone group (p < 0.0001). No significant differences in AEs were noted between groups [52].

NB-UVB in Combination with Biologic Therapies

A prospective clinical trial evaluated the efficacy of etanercept (ETN) in combination with NB-UVB in the treatment of psoriasis. In the cohort, NB-UVB had been administered as a first-line treatment to 322 subjects with moderate-to-severe psoriasis. Subjects who were unresponsive to NB-UVB (defined as failing to achieve a PASI75 at 8 weeks) were evaluated for treatment with conventional systemic therapy (MTX, cyclosporine, acitretin, PUVA). If subjects were ineligible for conventional therapies or failed to respond, they were treated with ETN 50 mg twice weekly. If subjects subsequently failed to achieve PASI75 in 12 weeks on ETN alone, they were treated with NB-UVB plus ETN. A total of eight patients met these criteria. Of the eight that received NB-UVB plus ETN, all achieved a PASI75 after 14.6 ± 3.3 exposures to NB-UVB. No significant AEs were observed in patients receiving this combination [53].

In a study assessing the efficacy of ETN plus NB-UVB, 99 subjects received a 12-week course of ETN. Seventy-five subjects did not achieve PASI90 after 12 weeks of ETN alone and were randomized to receive etanercept 50 mg once weekly monotherapy or in combination with NB-UVB three times weekly over a 4-week time period. Of the subjects with high adherence to NB-UVB, PASI90 was achieved at week 16 in 42.9% of subjects in the ETN plus NB-UVB group compared to 3.4% in the ETN monotherapy group (p = 0.018). There were no significant differences in AEs in either group [54].

The combination of NB-UVB + ETN versus ETN alone was evaluated in 30 subjects with a BMI > 30. All subjects initially received 50 mg subcutaneous ETN twice weekly for 12 weeks. Subjects were then randomized to receive either a maintenance dose of 50 mg ETN weekly or NB-UVB plus 50 mg ETN maintenance dose. The NB-UVB plus ETN group had similar PASI75 scores at 24 weeks when compared to the ETN monotherapy group (53.3 vs. 46.7%) [55].

Targeted UVB

A recent study evaluated the use of a single dose of targeted UVB at ten times the MED (MED10) in 18 subjects with plaque psoriasis, excluding the face, hands, feet, scalp, or intertriginous areas. Subjects were evaluated 4 and 8 weeks after receiving a single MED10 dose. PASI scores were reduced from a baseline of 16.2 at treatment start, to 8.0 and 7.2 at 4 and 8 weeks, respectively (p = 0.005). AEs typically resolved within 3 days, were reported as mild, and included erythema, edema, and clear exudate at treated sites 1 to 2 days post treatment in 5 of the 18 subjects [56].

Excimer Laser

Excimer Laser Monotherapy

In a separate study evaluating the efficacy and safety of a 308nm excimer laser in the treatment of scalp and palmoplantar psoriasis, 41 subjects with resistant scalp or palmoplantar psoriasis were recruited. Twice-weekly treatment was initiated at three times the MED (MED3) and increased at intervals no greater than 20% for each subsequent visit. Single treatments were skipped if subjects exhibited sunburn-like reactions from the previous treatment cycle. Subjects with scalp psoriasis were evaluated at baseline using the Psoriasis Severity Scalp Index (PSSI) and re-evaluated at 2-week intervals until week 12. At 12 weeks, 13 subjects achieved at least a 75% reduction in PSSI, and 8 subjects achieved at least a 50% reduction in PSSI. In the palmoplantar psoriasis group, 15 subjects were evaluated at baseline using the Psoriasis Severity Index (PSI). PSI scores at baseline were 7.4 and were reduced to 1.3 after 12 weeks of laser treatment. Clinical response was observed in as few as one or two treatments. Treatment was well tolerated; side effects generally consisted of sunburn-like reactions such as hyperpigmentation, erythema, crusting, blistering, or itching [57].

Excimer Laser in Combination with Topical Therapies

A 12-week, open-label, pilot study assessed the efficacy of calcitriol 3 µg/g ointment and clobetasol dipropionate 0.05% spray in combination with the XTRAC® 308 nm excimer laser in the treatment of psoriasis. Twenty-one subjects with \leq 10% BSA plaque psoriasis were recruited. All subjects were treated with excimer laser twice weekly for the first 6 weeks. Subjects who did not achieve PASI75 after week 6 were treated with the excimer laser twice weekly as needed for weeks 7 through 12. Dosing was based on induration of the target lesion. Topical treatment was rotated during the 12-week study period in combination with excimer laser. During weeks 1 through 4, subjects were treated with clobetasol spray twice daily. During weeks 5 through 8, subjects were treated with calcitriol ointment twice daily. During weeks 9 through 12, subjects were treated with clobetasol spray and calcitriol ointment twice daily. At week 12, 76% of subjects achieved PASI75, and 52% of subjects achieved a PGA score of clear or almost clear. AEs were mild and consisted of short-term phototoxic reactions [58].

UVA and PUVA

PUVA treatment can be administered using oral 8methoxypsoralen, or via warm bath administration in which psoralen derivatives have been dissolved. A two-armed multicenter trial evaluating the two approaches was undertaken in 74 subjects with moderate-to-severe psoriasis. No significant difference in efficacy was observed in the two treatment groups. Bath PUVA may be an effective option for patients with contraindications to receiving oral PUVA [59].

In a study comparing the efficacy of BB-UVA and PUVA in the treatment of psoriasis, 61 subjects with > 30% BSA involvement were enrolled. Thirty subjects received PUVA, 16 received BB-UVA at 10 J/cm², and 15 received BB-UVA at 15 J/cm². The BB-UVA group achieved similar results until session 24, but then plateaued, while the PUVA group continued to achieve improvement. The study concluded at 48 sessions. Phototoxic reactions were the most common side effect and occurred more frequently and with greater severity in the BB-UVA 15-J/cm² group [60]. The efficacy of UVA1, which uses non-erythemogenic wavelengths between 340 and 400 nm, was evaluated in the treatment of 62 subjects with palmoplantar pustulosis (PPP). Subjects were evaluated using the palmoplantar pustular psoriasis area and severity index (PPPASI) score. Subjects with a baseline score of 9.4 were treated for up to 30 sessions with UVA1 light three times per week with a wavelength range of 340 to 400 nm (peak emission at 365 nm). A 75% reduction in PPPASI (1.7 ± 1.9) score was achieved in 72.6% of subjects by session 30. AEs included hyperpigmentation, pruritus, and sunburn-like reactions [61].

Recent studies have evaluated the efficacy of oral psoralen plus sunlight (PUVAsol) [62, 63]. In a randomized prospective clinical trial comparing the clinical and cost effectiveness of PUVAsol versus PUVA therapy in the management of psoriasis, subjects were randomized to receive either PUVA treatment or PUVAsol treatment. Thirty-six subjects completed the study. The reduction in PASI score was greater in the PUVA group at 2 and 4 weeks compared to the PUVAsol group (p =0.034). At 8 weeks, 12 weeks, and endpoint, treatment response was comparable in both groups [62]. In a study assessing the efficacy of PUVAsol versus the combination of oral isotretinoin and PUVAsol, 40 subjects with psoriasis were randomized to receive PUVAsol alone or PUVAsol plus oral isotretinoin. Subjects who received PUVAsol plus oral isotretinoin achieved PASI75 in a mean of 8 weeks while subjects who received PUVAsol alone achieved PASI75 in a mean of 10 weeks [63]. Variability in the dose of UV light is inevitable with use of PUVAsol, but PUVAsol has a similar efficacy and side effect profile as traditional PUVA and may be beneficial in resource-limited countries.

LED Light

The use of LED light therapy in the treatment of psoriasis has emerged as a potential treatment option [64]. LED blue light (LED-BL) with peak wavelengths ranging from 420 to 453 nm is efficacious and associated with minor side effects such as hyperpigmentation [65, 66].

The efficacy of 630-nm peak wavelength LED red light (LED-RL) compared to 430-nm peak LED-BL was evaluated in 20 subjects with psoriasis. Clinical improvement was noted in both treatment groups. Improved erythema of psoriatic plaques was noted throughout the study period in those who receive LED-BL, while those who received LED-RL had no improvement in erythema beyond six illuminations. Patient-reported hyperpigmentation was less in subjects who received LED-RL [67].

The efficacy of LEDs emitting in the UVB spectrum (UVB-LED) was evaluated in a prospective, right-left comparative, open study. Twenty subjects with psoriasis were enrolled; symmetrical psoriasis lesions on the extremities or

trunk were chosen. One lesion was treated with UVB-LED with aggressive dose escalation or slow dose escalation; the other lesion was left untreated and served as control. Overall improvement at the end of treatment was 93% in the aggressive dose escalation group and 84% in the slow dose escalation group. Results were similar to NB-UVB delivery by traditional bulbs, providing evidence that a conversion to more cost-effective LED lighting may be an option pending the availability of more evidence to support this switch [68].

Conclusions

Barriers to treatment with phototherapy exist and compliance may be a challenge for patients [3]. The use of phototherapy may be impeded by geographic accessibility of phototherapy units, cost of travel, and personal and professional conflicts that interfere with consistent treatment [69••]. Home phototherapy options may help address these hurdles. Phototherapy remains a safe and efficacious treatment modality in psoriasis patients with optimal treatment adherence.

There has been a rapid expansion in the available treatment options for psoriasis. New pharmacologic therapies which target specific immune pathways have emerged. However, clinical trials assessing the role of phototherapy in combination with emerging biologic therapies are lacking. Limited data suggests that simultaneous use of phototherapy modalities and biologic therapies may be safe and beneficial in select patients.

Approximately 10–15% of patients with psoriasis receive some form of light therapy annually [70••]. Phototherapy should continue to be a first-line treatment for patients with moderate-to-severe disease or in those who are unresponsive to topical treatment. Future studies should evaluate the use of phototherapy in comparison to and in conjunction with recent and emerging treatment options.

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Compliance with Ethical Standards

Conflict of Interest Feldman has received research, speaking, and/or consulting support from a variety of companies including Galderma, GSK/Stiefel, Almirall, Leo Pharma, Baxter, Boeringer Ingelheim, Mylan, Celgene, Pfizer, Valeant, Taro, Abbvie, Cosmederm, Anacor, Astellas, Janssen, Lilly, Merck, Merz, Novartis, Regeneron, Sanofi, Novan, Parion, Qurient, National Biological Corporation, Caremark, Advance Medical, Sun Pharma, Suncare Research, Informa, UpToDate, and National Psoriasis Foundation. He is the founder and majority owner of www.DrScore.com and the founder and part owner of Causa Research, a company dedicated to enhancing patients' adherence to treatment. Seth Howell and Leah Cardwell have no conflicts to disclose.

Human and Animal Rights and Informed Consent This article does not contain any studies with human or animal subjects performed by any of the authors.

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