

# Evidence and Considerations in the Application of Chemical Peels in Skin Disorders and Aesthetic Resurfacing

<sup>a</sup>MARTA I. RENDON, MD; <sup>b</sup>DIANE S. BERSON, MD, FAAD; <sup>c</sup>JOEL L. COHEN, MD, FAAD;  
<sup>d</sup>WENDY E. ROBERTS, MD; <sup>e</sup>ISAAC STARKER, MD, FACS; <sup>f</sup>BEATRICE WANG, MD, FRCP, FAAD

<sup>a</sup>Clinical Associate Professor, Dermatology, University of Miami School of Medicine, Miami, Florida; <sup>b</sup>Assistant Professor, Dermatology; Assistant Attending Physician, New York-Presbyterian Hospital, New York, New York; <sup>c</sup>Director, AboutSkin Dermatology and DermSurgery, PC; Clinical Associate Professor, Dermatology, University of Colorado, Denver, Colorado; <sup>d</sup>Assistant Clinical Professor of Medicine, Loma Linda University Medical Center, Loma Linda, California; <sup>e</sup>Clinical Private Practice, the Peer Group Plastic Surgery Center, Florham Park, New Jersey; <sup>f</sup>Director, Melanoma Clinic; Assistant Professor, McGill University, Montreal, Canada

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## ABSTRACT

Chemical peeling is a popular, relatively inexpensive, and generally safe method for treatment of some skin disorders and to refresh and rejuvenate skin. This article focuses on chemical peels and their use in routine clinical practice. Chemical peels are classified by the depth of action into superficial, medium, and deep peels. The depth of the peel is correlated with clinical changes, with the greatest change achieved by deep peels. However, the depth is also associated with longer healing times and the potential for complications. A wide variety of peels are available, utilizing various topical agents and concentrations, including a recent salicylic acid derivative,  $\beta$ -lipohydroxy acid, which has properties that may expand the clinical use of peels. Superficial peels, penetrating only the epidermis, can be used to enhance treatment for a variety of conditions, including acne, melasma, dyschromias, photodamage, and actinic keratoses. Medium-depth peels, penetrating to the papillary dermis, may be used for dyschromia, multiple solar keratoses, superficial scars, and pigmentary disorders. Deep peels, affecting reticular dermis, may be used for severe photoaging, deep wrinkles, or scars. Peels can be combined with other in-office facial resurfacing techniques to optimize outcomes and enhance patient satisfaction and allow clinicians to tailor the treatment to individual patient needs. Successful outcomes are based on a careful patient selection as well as appropriate use of specific peeling agents. Used properly, the chemical peel has the potential to fill an important therapeutic need in the dermatologist's and plastic surgeon's armamentarium.

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Chemical peels are used to create an injury of a specific skin depth with the goal of stimulating new skin growth and improving surface texture and appearance. The exfoliative effect of chemical peels stimulates new epidermal growth and collagen with more evenly distributed melanin. Chemical peels are classified by the depth of action into superficial, medium, and deep peels.<sup>1</sup> Specific peeling agents should be selected based on the disorder to be treated and used with an appropriate peel depth, determined by the histological level or severity

of skin pathology to maximize success. However, other considerations, such as skin characteristics, area of skin to be treated, safety issues, healing time, and patient adherence, should also be taken into account for best overall results.

Chemical peels are very common in clinical practice. The American Society of Plastic Surgery reported that more than one million peel procedures were performed by its members in 2008.<sup>2</sup> Although peels have recently had an upsurge in research interest,<sup>3</sup> they are best

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**ADDRESS CORRESPONDENCE TO:** Marta I. Rendon, MD, The Dermatology and Aesthetic Center, 880 NW 13th Street, #3C, Boca Raton FL 33486; E-mail: skincareresearch@bellsouth.net

**TABLE 1. Action of peeling molecules in the skin**

ULTRASTRUCTURE	LHA		SALICYLIC ACID		LACTIC ACID	
	1%	5%	3%	15%	3%	15%
Dermo epidermal junction—separation	+	+++	+++	+++	0	0
Alive epidermis						
Expanded spaces	0	0	+++	+++	0/+	0/+
Vacuoles	0	0	+++	+++	0	0
Perinuclear edema	+	+	+++	+++	+	+
Spoiled membranes	0	0	+++	+++	0	0
Spoiled desmosomes	0	0	+	++	0	0
Granular junction	0	0	+	+++	0	++
Spoiled corneodesmosomes	0	0	++	+++	0/+	+
Stratum compactum—basket weave pattern (BWP)	0	0	+++	+++	++	+++
Central spoiled corneosomes	0	0	+++ (D)	+++ (D)	++	+++
Peripheral spoiled corneosomes	0	0	++	+++	0	0
Spoiled proteic cornea envelope	0	0	+	++	0	0
Spoiled lipidic cornea envelope	0	0	+	++	0	0
Stratum junction compactum/stratum disjunctum BWP	0	+	Nonidentifiable			
Central spoiled corneosomes—rupture	+++	+++				
Peripheral spoiled corneosomes—rupture	0	0				
Spoiled cornea envelope	+	++				
Saw tooth corneocytes	+	++				
Stratum disjunctum BWP	+	+++	+++	+++	+++	+++
Peripheral spoiled corneosomes—rupture	0	+++	+++	+++ (D)	++	+++
Spoiled proteic cornea envelope	+	++	+++	+++	+++	+++
Spoiled lipidic cornea envelope	+	+++	+++	+++	+++	+++
Saw tooth corneocytes	+	+++	+++	+++	+++	+++
Keratin content	0	0	0	0	0	0

Adapted from Berson D, et al. *J Drugs Dermatol*. 2009;8:803–811

performed and/or supervised by dermatologists and plastic surgeons who have far more experience and knowledge with cosmetic procedures than other physicians.<sup>3</sup>

Using the correct depth chemical peel is a critical component for success. Superficial peels affect the epidermis and dermal-epidermal interface. They are useful in the treatment of mild dyschromias, acne, post-inflammatory pigmentation, and AKs and help in achieving skin radiance and luminosity. Because of their superficial action, superficial peels can be used in nearly all skin types. After a superficial peel, epidermal regeneration can be expected within 3 to 5 days, and desquamation is usually well accepted. Superficial peels exert their actions by decreasing corneocyte adhesion and increasing dermal collagen.<sup>1</sup> These peels are a good method for rejuvenating the epidermis and upper dermal layers of skin.

Medium-depth peels may be used in the treatment of dyschromias, such as solar lentigines, multiple keratoses, superficial scars, pigmentary disorders, and textural changes. The healing process is longer, with full epithelialization occurring in about one week. Sun protection after a medium-depth peel is recommended for several weeks. Because of the risk of prolonged hyperpigmentation, medium-depth peels should be conducted with caution in patients with dark skin.

Deep peels may be used for severe photoaging, deep or coarse wrinkles, scars, and sometimes precancerous skin lesions. Usually performed with phenol in combination with croton oil, deep peels cause rapid denaturation of surface keratin and other proteins in the dermis and outer dermis. Penetrating the reticular dermis, the deep peel maximizes the regeneration of new collagen. Epithelialization occurs in 5 to 10 days, but deep peels require significant healing time, usually two months or more, and sun protection must always be used. Phenol is rapidly absorbed into the circulation, potentiating cardiotoxicity in the form of arrhythmias. Therefore, special care, such as cardiopulmonary monitoring and intravenous hydration, must be provided to address this concern.<sup>4,5</sup> Other complications include hypopigmentation, hyperpigmentation, scarring, and keloid formation, which may occur primarily with phenol peels (similar to laser resurfacing, the occurrence of these problems is both operator- and technique-dependent).<sup>6</sup> Phenol peels are primarily performed in operating room settings and are frequently used as adjuncts to surgical procedures. Due to the increased risk of prolonged or permanent pigmentary changes, deep peels are not recommended for most dark-skinned individuals. Currently, new laser techniques are a popular alternative for major deep skin resurfacing because they avoid the adverse effects of deep chemical peels, even if phenol is used in lower concentrations.

Chemical peels are a mainstay in the cosmetic practitioner's armamentarium because they can be used to treat some skin disorders and can provide an aesthetic benefit. In addition, chemical peels may be readily combined with other resurfacing and rejuvenation procedures, often providing synergistic treatment and

more flexibility in tailoring treatments to specific patient needs and conditions. Clinicians can customize regimens to the patient's individual needs using several modalities, such as at-home skin regimens, chemical peels, and lasers or dermabrasion, to provide unheralded flexibility in individualized care.

This brief review covers chemical peels and their role in appropriate indications by combining evidence-based medicine with the clinical experience of the authors. The recent introduction of  $\beta$ -lipohydroxy acid, a salicylic acid derivative with antibacterial, anti-inflammatory, antifungal, and anticomedogenic properties, may provide additional therapeutic benefit, and thus its role is highlighted.

## CURRENTLY AVAILABLE PEELS

A wide variety of peels are available with different mechanisms of actions, which can be modulated by altering concentrations. Agents for superficial peels today include the alpha hydroxy acids (AHAs), such as glycolic acid (GA), and the beta hydroxy acids (BHAs), including salicylic acid (SA). A derivative of SA,  $\beta$ -lipohydroxy acid (LHA, up to 10%) is widely used in Europe and was recently introduced in the United States. Tretinoin peels are used to treat melasma and postinflammatory hyperpigmentation (PIH).<sup>7</sup> Trichloroacetic acid (TCA) can be used for superficial (10–20%) peels and for medium-depth peels (35%). Combination peels, such as Monheit's combination (Jessner's solution with TCA),<sup>8</sup> Brody's combination (solid carbon dioxide with TCA),<sup>9</sup> Coleman's combination (GA 70% + TCA),<sup>10</sup> and Jessner's solution with GA,<sup>11</sup> have been used for medium-depth peels where a deeper effect on the skin is required but deep peeling is not an option. Deep peels are typically performed with phenol-based solutions, including Baker-Gordon phenol peel and the more recent Hetter phenol-croton oil peel.<sup>12</sup>

The recent introduction of LHA is important because it not only provides efficient exfoliation at low concentrations, it possesses antibacterial, anti-inflammatory, antifungal, and anticomedogenic properties.<sup>13–15</sup> An SA derivative with an additional fatty chain, LHA has increased lipophilicity compared to SA, for a more targeted mechanism of action and greater keratolytic effect.<sup>13</sup> LHA has good penetration into the sebaceous follicle and through the epidermis, but it penetrates less deeply into the skin than GA or SA (LaRoche-Posay; data on file; 2008) interacting with the more superficial layers of the stratum corneum, specifically the compactum/disjunctum interface. Thus, its activity focuses on the follicle and epidermis. LHA has a pH similar to normal skin (pH 5.5) and has proven to be quite tolerable. Conveniently, the LHA peel does not require neutralization in contrast to a GA peel.

LHA has an interesting mechanism of action. It targets the corneosome/corneocyte interface to cleanly detach individual corneosomes, which may partially explain skin smoothness after an LHA peel, since it minimizes desquamation of clumps, which leads to roughness.<sup>14</sup> These effects are visible to the naked eye.<sup>13</sup> Similar to SA, LHA

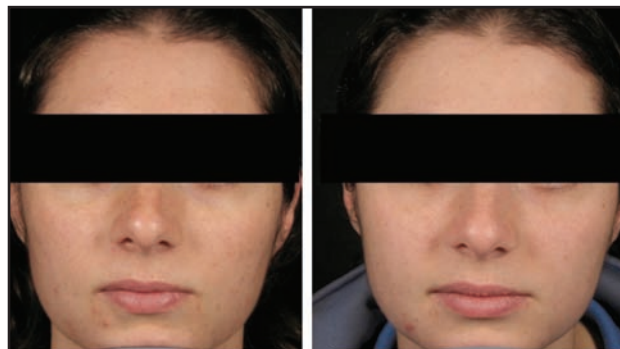
does not affect keratin fibers or the corneocyte membrane.<sup>13</sup> AHAs and BHAs do not modify corneocyte keratins. The clean and uniform corneocyte separation achieved with LHA more closely mimics the natural turnover of skin. SA and GA can result in only partial detachment of some cells, which leads to uneven exfoliation of cells in clumps. The differences between LHA, SA, and lactic acid with regard to epidermal effects are summarized in Table 1. The histological section of skin samples treated with LHA also shows targeting of the horny layer by LHA along with good epidermal integrity. Studies have demonstrated that LHA targets corneodesmosome protein structures, particularly corneodesmosine, in the horny layer (LaRoche-Posay; data on file; 2008). While SA has the same target, its activity is less specific and is limited to arbitrary intercellular cleaving of some intercellular junctions. Finally, AHAs have far less affinity for these proteins and the less drastic cleaving of the intercellular bonds of SA leads to less precise desquamation than that observed with LHA.

Other properties of LHA include modifying the stratum corneum so that postpeel, it is thinner, flexible, and resistant to wrinkling and cracking.<sup>16</sup> *In-vivo* immunohistological study of LHA peels showed increased epidermal thickness and dendritic hyperplasia without markers of irritation or inflammation.<sup>15</sup> Thus, LHA has similar effects to those of SA on epidermal indices, such as thickness of stratum corneum and germinative compartment and number of nuclei.<sup>14,17</sup> Additionally, LHA-treated older skin has been shown to recover some physiological characteristics of younger skin, such as more rapid cell cycling.<sup>14</sup> LHA has very few side effects. In clinical studies, LHA peels were well tolerated with some patients experiencing burning and crusting after the initial peel. No cases of PIH or scarring have been reported with LHA.<sup>18</sup>

## APPLICATIONS OF PEELS IN CLINICAL PRACTICE

**Acne.** Clinicians and patients often use chemical peels as an adjunct to medical therapy in acne because they produce complementary rapid therapeutic effects and improvements in skin appearance and textures.<sup>19,20</sup> The primary effect may be on comedones with a concomitant reduction in inflammatory lesions (Figures 1–3). Peels may allow topical acne agents to penetrate more efficiently into the skin and may improve PIH.<sup>21</sup> With good technique, peels may also be beneficial for dark-skinned patients who have pigmentary changes due to acne.<sup>20</sup> While 2009 American Academy of Dermatology guidelines suggest that more evidence is needed to determine best practices,<sup>22</sup> clinical experience has shown promising utility. Peels that have been studied for active acne include SA, GA, LHA, and Jessner's solution.

**SA.** SA can be used to treat comedones and inflammatory lesions.<sup>21</sup> In the early 1980s, a controlled, double-blind trial (N=49) showed that low concentrations of SA (0.5–3%) helped speed resolution of inflammatory lesions.<sup>23</sup> Later, Lee et al<sup>18</sup> reported improvement in acne in 35 Korean patients with acne treated with SA 30% peels, and that the reduction in lesion counts increased as



**Figure 1.** Patient with mild inflammatory acne before and after LHA peeling shown before (left) and after four sessions at two-week intervals (right). Photo courtesy of Joel L. Cohen, MD.

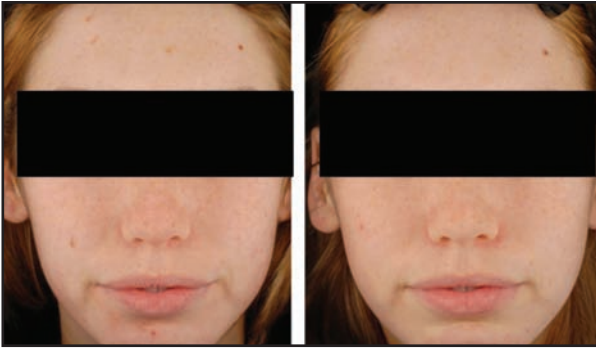
the duration of peel continued.<sup>18</sup> SA has shown good effects in dark-skinned Asian, African-American, and Hispanic patients with acne.<sup>24,25</sup> In addition, this treatment regimen facilitated resolution of PIH as well as a decrease in the overall pigmentation of the face.<sup>25</sup>

Most recently, Kessler et al<sup>26</sup> compared 30% GA versus 30% SA peels in 20 patients with mild-to-moderate acne using a split-face design. Peels were performed every two weeks for a total of six treatments. Both peels improved acne; however, the authors found that the SA peel had better sustained efficacy (number of acne lesions, improvement rating by blinded evaluator) and fewer side effects than GA, presumably due to the increased lipophilicity of SA.<sup>26</sup> Overall, the authors of this paper agree with the impression that SA peels are better tolerated than GA peels in acne patients.

**LHA.** Due to its lipophilicity, LHA targets the sebum-rich pilosebaceous units and has a strong comedolytic effect. Uhoda et al<sup>27</sup> studied LHA in acne-prone women and women with comedonal acne (n=28) in a randomized, controlled, clinical trial. As shown with ultraviolet (UV) light video recordings and computerized image analysis, both the number and size of microcomedones were significantly decreased in 10 of 12 LHA-treated patients versus 3 of 10 untreated controls. In addition, image analysis showed a marked reduction in the density of follicular keratotic plugs. As microcomedones resolved, there was also a decrease in follicular bacterial load. There were no reported side effects with LHA use.<sup>27</sup>

The previously described anticomedogenic properties of LHA include loosening of both intercorneocyte binding and bacterial adhesion inside the follicular openings<sup>16</sup> and thinning of the stratum corneum.<sup>28</sup> LHA reduced the bacterial population per volume of follicular cast by 21±13 percent following daily treatment with a 2% cream. In addition, bacterial viability was reduced.<sup>28,16</sup>

**GA.** GA may be used in acne to normalize keratinization and increase epidermal and dermal hyaluronic acid and collagen gene expression.<sup>29</sup> It has been studied in concentrations ranging from 35 to 70%.<sup>19,30,31</sup> GA 70% has been shown to reduce comedones in Asian patients.<sup>19</sup> Lower concentrations (35% or 50%)



**Figure 2.** Patient with mild inflammatory acne treated with LHA peels shown before (left) and after four sessions at two-week intervals (right). Photo courtesy of Marta Rendon, MD.



**Figure 3.** Patient with inflammatory acne and postinflammatory hyperpigmentation shown before (left) and after four LHA peeling sessions at two-week intervals (right). Photo courtesy of Marta Rendon, MD.

also achieved significant resolution of both inflammatory and non-inflammatory acne lesions.<sup>30</sup> Another study also conducted on Asian patients showed improvement in pigmentation problems and reported that acne flares after the first treatment diminished with subsequent treatments.<sup>30</sup> A case series suggested that comedones may improve more readily than inflammatory lesions,<sup>31</sup> but this remains to be validated.

**Jessner's solution.** Superficial Jessner's solution peels have been used to manage acne. Medium-depth peels involving Jessner's solution plus TCA have also been used to treat mild acne scarring. Kim et al<sup>19</sup> compared Jessner's solution versus GA 70% in patients with facial acne in a split-face study (n=26). Efficacy was similar between the two types of peels, but Jessner's solution was associated with a significantly greater degree of exfoliation compared with GA ( $P<0.01$ ).<sup>19</sup> Lee et al<sup>32</sup> studied the effect of GA and Jessner's solution on facial sebum secretion in patients with acne.<sup>32</sup> GA 30% or Jessner's solution peels were performed twice at an interval of two weeks in 38 patients (27% GA, 11% Jessner's solution), and sebum levels were measured. In this study, neither type of peel changed sebum secretion after two peels.<sup>32</sup> However, Jessner's solution may be an option for superficial peeling as an adjunctive treatment in patients with acne.

**Acne scarring.** Acne scars are polymorphic; therefore, it is important to assess and design treatment according to the types of scars, while also keeping in mind patient expectations. Chemical peels, laser resurfacing, dermabrasion, and fractionated laser technology as well as fillers and subcision are commonly used modalities for acne scar therapy. From a peel standpoint, patients with mild-to-moderate acne scarring may be treated. Peels that have been used include SA, GA, TCA, LHA, and Jessner's solution. Peels are used as an adjunct to medical therapy including a retinoid or AHAs.<sup>33</sup> Studies of Jessner's solution in combination with TCA in medium-depth peels have also shown benefit in acne scarring.<sup>34,35</sup> Medium-depth and deep-depth phenol peels, while useful for treatment of acne scarring, are not recommended for dark skin types IV to VI due to a high risk of permanent pigmentary changes.<sup>36</sup> Regional dermabrasion is an effective adjunct to chemical peel for medium-depth scars.<sup>37</sup>

**Phenol solutions.** Deep chemical peels may be used to treat acne scarring. The most common solutions are combinations of phenol and croton oil.<sup>12,38-40</sup> These solutions penetrate to the midreticular region and maximize the production of collagen.<sup>41</sup> Park et al<sup>42</sup> used a modified phenol peel, which was applied to 46 patients of Asian descent, 11 of whom were treated for acne scarring and 28 for wrinkles. Seven of 11 patients (64%) with acne scars improved 51 percent or more based on physician and patient assessment. The most frequent side effect was PIH (74%).<sup>42</sup>

**Photodamage.** Photodamaged skin is associated with chronic UV light exposure. Photoaging changes include a thicker dermis due to breakdown of the elastic fiber network and a thinner epidermis having cellular atypia. Often, the result can be irregular pigmentation, wrinkling, loss of elasticity, development of solar lentigines and actinic keratoses, and coarseness. Histologically, peels alter the epidermis creating a more normal pattern with columnar cells showing return of polarity, more regular distribution of melanocytes, and melanin granules. A wide range of chemical peels including AHA, SA, TCA, and phenol are used to treat photodamage; selection is based on patient presentation and severity of photodamage. The efficacy of treating photoaging with tretinoin is well established.<sup>16</sup> Efficacy of peels to treat photodamage has also been repeatedly reported. In photodamaged skin, peels cause skin exfoliation and rejuvenation,<sup>43</sup> and repeated superficial peels may be used.<sup>44</sup> With advanced photoaging changes, a peel may be combined with laser resurfacing or other procedures.

AKs are precancerous lesions that are also a result of chronic UV exposure. Peels have been used to treat AKs and are appropriate treatment for most regions of the body. Chemical peels can eliminate AKs and may be able to provide prophylaxis for a prolonged time period.<sup>45</sup> They have also recently shown clinical benefit when AKs were observed in combination with Bowen's Disease.<sup>46</sup>

SA. Kligman et al<sup>47</sup> studied SA 30% in regimens of

single and multiple peels at four-week intervals and reported improvement of pigmentation, skin texture, and reduction of fine lines in patients with moderately photodamaged skin. Humphreys et al<sup>48</sup> reported that 40% TCA (a borderline medium-depth peel) plus topical retinoid treatment improved solar lentigines, AKs, and skin texture, but had minimal effect on wrinkles.

GA. Rendon et al<sup>49</sup> described the use of superficial GA peels in combination with dermal fillers and botulinum toxin, successfully addressing wrinkles, uneven skin tone, skin laxity, and skin clarity. They used a schedule that separates fillers and peels by approximately one week; with botulinum toxin, the peel was administered after the toxin in the same visit or the procedures were separated by one or more days to minimize the potential for side effects.<sup>49</sup> Briden et al<sup>50</sup> reported good patient satisfaction when using superficial GA peels with microdermabrasion in photoaging.

LHA. Efficacy of LHA peeling in photodamage was shown in a randomized, intraindividual-controlled, split-face trial evaluating LHA (5–10%) versus GA peel (20–50%) (LaRoche-Posay; data on file; 2008). A total of 43 women with fine lines, wrinkles, and hyperpigmentation were treated with six applications with both acids over nine weeks. Both treatments showed a significant effect in reducing fine lines, wrinkles, and hyperpigmentation (Figure 4). However, the efficacy of four LHA sessions was equivalent to six sessions of GA. The LHA peel was well tolerated. No patient withdrew from the study, and the most common side effect was transient erythema that persisted for less than two hours (LaRoche-Posay; data on file; 2008).

Leveque et al<sup>14</sup> assessed skin improvement in 80 women who were treated with an excipient containing LHA 1% daily for six months, finding a progressive improvement in complexion, with an onset of action occurring within one month. In a randomized, controlled trial comparing GA 10% versus LHA 2% versus retinoic acid 0.05% on the forearm, LHA and retinoic acid improved surface texture similarly while GA had a very minimal effect.<sup>51</sup>

AHA increases UV sensitivity,<sup>52,53</sup> while LHA increases the skin's resistance to UV-induced damage. Saint-Leger<sup>16</sup> reported that the minimal erythema dose was 210mJ/cm<sup>2</sup> versus 140mJ/cm<sup>2</sup> for untreated and placebo-treated controls (LaRoche-Posay; data on file; 2008).<sup>16</sup> This protective effect may be due to the antioxidant properties of LHA, which can inactivate the oxygen singlet (<sup>1</sup>O<sub>2</sub>) without reacting with it and thus quench the superoxide anion. It also reacts avidly with hydroxyl radicals to produce 2,5-dihydrobenzoic acid, an excellent scavenger of the superoxide anion (L'Oreal; data on file; 2008).<sup>16</sup>

Combination solutions. Lawrence et al<sup>54</sup> conducted a 15-patient, split face study comparing a medium-depth chemical peel consisting of Jessner's solution and 35% TCA with topical fluorouracil in the treatment of widespread facial AKs. Both treatments reduced the number of visible AKs by 75 percent and produced equivalent reductions in



**Figure 4.** Patient with photoaging-related pigmentary changes shown before (top) and after four LHS peel sessions at two-week intervals (bottom). Photo courtesy of Marta Rendon, MD.

keratinocyte atypia, hyperkeratosis, parakeratosis, and inflammation, with no significant alteration of preexisting solar elastosis and telangiectasia.<sup>54</sup> Also, a 70% glycolic peel and a 5% 5-fluorouracil solution (Drogaderma, Sao Paulo, Brazil) was used in actinic porokeratosis every two weeks for four months with benefit, but the results remain to be validated.<sup>55</sup>

Phenol solutions. A study by Chew et al<sup>56</sup> suggested that there was a greater improvement in upper-lip wrinkles with Baker's phenol chemical peel than with CO<sub>2</sub> laser treatment ( $p < 0.03$ ), although the change from baseline was statistically significant for both chemical peel and CO<sub>2</sub> laser. In basal cell carcinoma, Kaminaka et al<sup>57</sup> demonstrated that nevoid basal cell carcinoma could successfully be treated with phenol and TCA peeling.<sup>57</sup> A more recent study by Kaminaka et al<sup>46</sup> not only demonstrated a significant benefit of the phenol-base peel in patients with AKs and Bowen's Disease, but also identified biomarkers that assisted in predicting clinical success from failure. They studied 46 patients treated with phenol peels and followed up for one or more years. Biopsy specimens were taken before and after treatment. In this small but important study, 39 patients (84.8%) had a complete response after 1 to 8 treatment sessions. Statistical differences also correlated the number of treatment sessions with histology, personal history of skin cancer, tumor thickness, and cyclin A expression. The authors concluded that tumor thickness and cyclin A could be specific and useful biomarkers as an accurate therapeutic diagnosis tool, thus providing a more useful way to measure potential therapeutic benefit.<sup>46</sup>

Melasma. Patients with melasma usually present with irregular patches of darkened skin on the cheeks, forehead, upper lip, nose, and chin.<sup>58</sup> Melasma has always been very challenging to treat for multiple reasons including the presence of melanin at varying depths in the

epidermis and dermis. Because chemical peels remove melanin and improve skin tone and texture, they are commonly used in treating this condition. More superficial and more limited involvement melasma is often more responsive to treatment. Data from small studies suggest that melasma improvement occurs more rapidly when peels are combined with medical therapy. Several peels have been studied (SA, LHA, GA, TCA, tretinoin and resorcinol, retinoic acid and Jessner's), although GA is currently most popular.

SA. Grimes<sup>25</sup> reported that a series of five SA peels at concentrations of 20 to 30% plus hydroquinone at two-week intervals resulted in moderate-to-significant improvement in 66 percent of six darker skinned (V–VI) patients. The treatment was well tolerated, and there was no residual hypo- or hyperpigmentation.<sup>25</sup> In unpublished data, Grimes noted that SA peels without hydroquinone preparation were associated with hyperpigmentation. Because of the known propensity of darker skin to develop dyschromias, Grimes recommended that even superficial peels be used with care and caution.

GA. In a study of GA 30 to 40% peels plus a modified Kligman's formula (retinoid, corticosteroid, and hydroquinone) versus Kligman's formula alone (n=40), Sarkar<sup>58</sup> found a significant decrease in Melasma Area and Severity Index (MASI) score from baseline to 21 weeks in both groups. Figure 5 shows an 80-percent change in score at Week 21 in the peel group and a 63-percent change in the control group ( $P<.001$ ).<sup>58</sup> However, the addition of a peel achieved a significantly greater effect versus the control group of Kligman's formula alone (more rapid and greater improvement,  $P<.001$ ).<sup>58</sup> Erbil et al<sup>61</sup> studied serial GA peels (from 35–50% and 70% every second peel) plus combination topical therapy (azelaic acid and adapalene) in 28 women with melasma<sup>59</sup> and found better results in the group receiving chemical peels plus topical therapy ( $P=0.048$ ), but only when the GA concentration was 50% or higher.<sup>59</sup> GA peels in concentrations of 20 to 70% administered every three weeks were studied alone or in combination with a topical regimen of hydroquinone plus 10% GA in 10 Asian women<sup>60</sup> in which the combination trended toward significance ( $P>0.059$ ).

In another study, a triple combination cream consisting of fluocinolone acetonide 0.01%, hydroquinone 4%, and tretinoin 0.05% was used in an alternating sequential treatment pattern, cycling with a series of GA peels, for the treatment of moderate-to-severe melasma.<sup>61</sup> Spectrometry measurements of the difference in melanin for involved versus uninvolved skin confirmed that hyperpigmentation was significantly reduced at Weeks 6 and 12 compared with baseline ( $P<0.001$  for both), with evaluations showing 90-percent improvement or more by Week 12 with the treatment approach.<sup>61</sup>

TCA. Kalla et al<sup>62</sup> compared 55 to 75% GA versus 10 to 15% TCA peels in 100 patients with recalcitrant melasma. They reported that both the time to response and degree

of response were more favorable with TCA compared with GA; however, relapse was more common in the TCA group (25 vs. 5.9% in the GA group).<sup>62</sup> Soliman et al<sup>63</sup> reported that 20% TCA peels plus topical 5% ascorbic acid was superior to TCA peeling alone in 30 women with epidermal melasma.

*Other peels.* An early report by Karam<sup>64</sup> used a 50% solution of resorcinol in patients with melasma and skin types I to IV.<sup>64</sup> A more recent study of 30 patients with mostly Fitzpatrick type IV skin type were treated successfully with lactic acid in a split-face comparison with Jessner's solution (N=30). All patients showed significant improvement as calculated by MASI score before and after treatment.<sup>65</sup> Khungar et al described a pilot study in which serial 1% tretinoin peels were as effective a therapy for melasma in dark-skinned individuals as 70% GA.<sup>7</sup>

**Potential side effects of peels.** Superficial peels are safe and tolerated with mild discomfort, such as transient burning, irritation, and erythema.<sup>66</sup> Scarring is rare in superficial peels, as are PIH and infection. In medium and deep peels, lines of demarcation that are technique related can occur. Care should be taken to feather peel solution at junctions with nonpeeled skin to avoid this effect. Side effects of deeper peels can also include pigmentary changes (e.g., PIH for dark-skinned individuals), infections, allergic reactions, improper healing, hypersensitivity, disease exacerbation, and those due to improper application.<sup>67,68</sup>

Care must also be taken to prophylactically treat patients with a history of herpes simplex infections. Herpetic episodes, usually on the lip or above the vermilion border, may be prevented with prophylactic oral acyclovir, valacyclovir hydrochloride, or famciclovir.<sup>69,70</sup> Antiviral agents are especially useful in patients who indicate a strong history of multiple herpetic lesions each year.

The best way to prevent complications is to identify patients at risk and maintain an appropriate peel depth that balances efficacy with known adverse events. Patients at risk include those with PIH, keloid formation, heavy occupational sun exposure, a history of intolerance to sunscreens, and uncooperative patients.

Tolerability of peels may be influenced by many factors, such as peel agents, concentration, depth, skin type, and concomitant use of skin care products. PIH can be exacerbated by sun exposure, so it is important to educate patients and closely monitor their recovery phase. Sunscreens should be used continuously to limit PIH development. Epidermal PIH responds well to various treatments, while dermal PIH remains problematic. Pretreatment with bleaching agents before beginning therapy with peels decreases the appearance of PIH. Treatment options include hydroquinone or kojic acid or other tyrosinase inhibitors.

In medium and deep peels, a common location of scarring is on the lower part of the face,<sup>71</sup> due perhaps to greater tissue movement or more aggressive treatment.

Other rare causes of scarring include infections and premature peeling, making post-peel monitoring an essential component of management. Delayed healing and persistent redness are early warning signs, and treatment with topical antibiotics and potent topical corticosteroids should be initiated as soon as possible to minimize scarring. Resistant scars may be treated with dermabrasion or pulsed dye laser followed by silicone sheeting therapy.

Acneiform eruptions may occur during or after peeling, presenting as erythematous follicular papules. These eruptions respond to oral antibiotics used in acne treatment. Discontinuation of oily skin preparations is also recommended.

Milia usually appear 2 to 4 months after peels in up to 20 percent of patients undergoing medium and deep peels and may be treated with extraction or electrosurgery.

Medium-depth peels are associated with most of the complications described above, though most can be managed successfully. Medium- and deep-depth peels should be used with great caution on skin types IV to VI. Toxicity, although rare, has been reported with resorcinol, SA, and phenol deep peels.<sup>72</sup>

## CONSIDERATIONS WITH ETHNIC SKIN

Indications for peeling in dark-skinned patients include treatment of dyschromia, PIH, acne, melasma, scarring, and pseudofolliculitis barbae. Clinicians should evaluate the Fitzpatrick skin type and ethnic background as part of the process of selecting whether a peel is an appropriate therapy and which peel is best suited for the individual patient.<sup>73</sup> Different ethnicities may respond unpredictably to chemical peeling regardless of skin phenotype. An individual patient history of PIH is very important to take into account. Hexsel et al<sup>74</sup> point out that Latin-Americans and Hispanics have a diverse range of skin phototypes and pigmentation and are prone to an increased incidence of melasma and PIH. In this subpopulation, they recommend peels as second-line therapy after topical therapies fail.<sup>74</sup>

Superficial peels may be safely used in patients with dark skin, including LHA 5 to 10%, TCA 10 to 20%, GA 20 to 70%, SA 20 to 30%, lactic acid, and Jessner's solution. In addition, variations of peel technique may be used, including spot treatment of PIH. This may be performed with TCA 25%, Jessner's solution, SA, and LHA. Table 2 provides recommended agents for peeling in dark-skinned individuals by specific indication. Deep phenol peels are not recommended for dark skin types IV to VI due to the high risk of prolonged or permanent pigmentary changes.<sup>75</sup> However, Fintsi et al<sup>76</sup> described safe use of phenol-based peels in patients with olive and dark skin and dark eyes and hair.<sup>76</sup>

## GENERAL APPROACH TO SKIN CARE BEFORE AND AFTER PEELING

**Medical history.** Taking a complete history prior to peeling is critical. It can enhance aesthetic results by

identifying any factors that may contribute to problems and provides an opportunity to discuss adherence issues necessary for successful management.<sup>67,77</sup> It is important to gain insight into patients' perceptions of wound healing and scar formation, as well as prior experience with resurfacing procedures or facelift surgery.<sup>67</sup> Current literature recommends waiting at least six months after discontinuing oral isotretinoin therapy before performing resurfacing procedures.<sup>67</sup>

A current medication list should be obtained, and photosensitizing agents should be discontinued. Some dermatological conditions, including rosacea, seborrheic or atopic dermatitis, and psoriasis, may increase the risk for postoperative problems, such as disease exacerbation, excessive and/or prolonged erythema, hypersensitivity, or delayed healing.<sup>67</sup> Prophylactic antiviral agents should be prescribed as required.<sup>67</sup> Since sun protection after peeling is essential, discussion in relation to the patient's past habits and experience is important.

**Pretreatment.** Pretreatment can help to enhance outcomes and is often started 2 to 4 weeks prior to the peel and discontinued 3 to 5 days before the procedure.<sup>21</sup> Topical retinoids or a prepeel solution can help to create a smooth stratum corneum to achieve a more even penetration of the peel. Topical retinoids may also speed healing.<sup>1</sup> Humphreys et al<sup>48</sup> reported that pretreatment with a topical retinoid resulted in more rapid and even frosting as well as a decrease in telangiectasias, which the authors postulated as being due to deeper penetration of TCA with retinoid pretreatment.<sup>48</sup>

Before a chemical peel, hydroquinone may be used to reduce the likelihood of PIH in dark-skinned individuals.<sup>1</sup> Discussing peel after-effects with patients before the peel is also important to aid comprehension of the peeling process.

Postpeel, patients should use a broad-spectrum sunscreen on a daily basis and implement a gentle cleansing regimen with toner and peel serum as prescribed. Moisturizers may also be recommended.

**Maintenance.** After a chemical peel, edema, erythema, and desquamation may occur for 1 to 3 days for superficial peels and 5 to 10 days for medium to deep peels. A cleansing agent may be used and antibacterial ointment applied especially for deep peels. Patients should be instructed to avoid peeling or scratching the affected skin and to use only simple moisturizers.

A long-term maintenance program will preserve the results of chemical peels in most patients. Patient participation and education is required, emphasizing the importance of sun protection and the use of appropriate skin care regimens that include cleansing, toning, exfoliation, and moisturizers. Patients need to have realistic expectations and understand that achieving benefits from peels requires repeated procedures. If the peel regimen works well for the patient, clinicians should consider a maintenance protocol, which may be one peel per month for six months, then every three months thereafter depending on the need and the season. Topical



**TABLE 2. Overview of chemical peels in dermatological conditions**

<b>ACNE</b>
<ul style="list-style-type: none"><li>• Chemical peels can be a useful adjunct to medical therapy for acne—may speed resolution, enhance penetration of topical drugs, and improve associated postinflammatory pigmentary problems</li></ul>
<ul style="list-style-type: none"><li>• Peels that have been studied for active acne include SA, LHA, GA 30–70%, and TCA 7–25%</li></ul>
<ul style="list-style-type: none"><li>• Need maintenance regimen—for patients who respond to peels, schedule every 3 months</li></ul>
<ul style="list-style-type: none"><li>• Primary effect may be on comedones, although reduction in inflammatory lesions may also occur (esp. SA, LHA)</li></ul>
<ul style="list-style-type: none"><li>• Peels also provide benefit in superficial acne scarring</li></ul>
<ul style="list-style-type: none"><li>• SA and its derivative LHA may often be the preferred peels for acne due to their action on both inflammatory and noninflammatory lesions</li></ul>
<b>PHOTODAMAGE</b>
<ul style="list-style-type: none"><li>• At least fair scientific evidence shows that the clinical benefits of chemical peels in photoaging outweigh the potential risks</li></ul>
<ul style="list-style-type: none"><li>• A range of chemical peels including AHAs, SA, LHA, TCA, and phenol are used to treat photodamage; selection is based on patient presentation (Glaugau, Fitzpatrick, PIH) and severity of photodamage</li></ul>
<ul style="list-style-type: none"><li>• SA, GA, TCA, phenol are all appropriate</li></ul>
<b>MELASMA</b>
<ul style="list-style-type: none"><li>• Peels are popular and widely used for melasma</li></ul>
<ul style="list-style-type: none"><li>• Peels may be most effective when used in combination with medical therapy or other procedures, possibly because peels remove melanin and other treatments inhibit melanocytes or melanogenesis</li></ul>
<ul style="list-style-type: none"><li>• Few formal studies are currently available; most existing studies have small patient populations</li></ul>
<ul style="list-style-type: none"><li>• Several peels have been studied in melasma (e.g., SA, LHA, GA, and TCA)</li></ul>
<ul style="list-style-type: none"><li>• Maintenance therapy is needed when peeling is used for melasma</li></ul>

retinoid maintenance therapy can also help maintain the skin rejuvenation results achieved with a chemical peel. It may be used alone on a daily or intermittent basis or in addition to 2 to 3 weekly light peels periodically. Maintenance regimens may also include products with combinations of kojic acid, hydroquinone, LHA, SA, GA, or ascorbic acid.

**Importance of tailoring therapy.** It is important to develop a peel program that is tailored to the individual needs of the patient. For example, a patient with visible photodamage who can tolerate social and work downtime may be treated with a 35% TCA peel while another patient may be better treated with a series of lighter peels

to minimize downtime. In addition, patients who are treated with peels may also be interested in a variety of other treatments, such as botulinum toxin or fillers, to improve the signs of aging.

## **CONCLUSION**

Chemical peels remain popular for the treatment of some skin disorders and for aesthetic improvement. Peels have been studied and shown to be effective as treatment for a myriad of conditions including acne, superficial scarring, photodamage, and melasma. Patients who are willing to undergo continued treatment are likely to be the best candidates. Newer molecules such as the LHA

superficial peel provide unique characteristics including targeted action and should be studied further. Clinicians should remember that there can be excellent synergy between peels and other procedures. Chemical peels are most effectively used in combination with a topical, at-home regimen, which, depending on the condition, may include exfoliating or moisturizing products, bleaching agents, or retinoids. Using peels less frequently but on a continuing basis is beneficial to help keep improvement ongoing, especially for superficial peels. Medium peels and deep peels are used more judiciously over time, but can address particularly difficult conditions effectively over the course of several treatments. Finally, it is important for patients to maintain a good sun protection regimen to optimize the clinical results achieved with chemical peels.

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