

Three-Dimensional Rejuvenation of the Photoaged Body

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Because of the multilayered pathophysiology of photoaging, a 1-dimensional approach targeting only a single skin layer is destined to fail. The treatment should address all affected skin layers in a multidimensional approach. Combining superficial and dermal procedures along with fillers for volume augmentation is the key to addressing these issues.

The 3-dimensional photoaging rejuvenation concept focuses on 3 types of rejuvenation corresponding to different skin layers. For each type, various scientifically proven procedures are available. However, combining the right techniques is sometimes difficult, as improvements in established modalities and new developments are continuously evolving. *Cosmet Dermatol.* 2011;24:381-387.

Photoaging is the premature aging of the skin caused by chronic UV exposure, particularly by UVA (320–400 nm) but also by UVB (290–320 nm) exposure. The clinical signs are discoloration, elastosis, deep wrinkles, and laxity, as well as inflammation and vascular lesions,¹ often accompanied by skin roughness and enlarged pores.² Furthermore, there also is a loss of volume attributable to lipoatrophy through transcriptional regulation of

lipogenic enzymes triggered by UV radiation.³ These factors together give photoaged skin its typical uneven and leathery appearance. In addition to the face, the most affected areas are the neck, décolletage, shoulders, forearms, and hands. Photoaging of the whole body is common in tanning bed users and sun worshippers.

Because of the multilayered pathophysiology of photoaging, a 1-dimensional approach targeting only a single skin layer is destined to fail. For a successful and satisfying outcome for the patient, the treatment needs to address all affected skin layers in a multidimensional approach. Combining superficial and dermal procedures along with fillers for volume augmentation is the key to addressing these issues. Although superficial treatments focus on discoloration and rough skin, dermal treatments focus on wrinkles, laxity, and vascular lesions. The deep dermal and subcutaneous approach is to tighten skin and replenish volume loss. For maximal safety and minimal

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downtime, rejuvenation approaches to the photoaged body should be nonablative, which corresponds to patients' expectations about the new natural look.⁴

The 3-dimensional photoaging rejuvenation concept focuses on 3 types of rejuvenation corresponding to different skin layers. The goals of type 1 rejuvenation are optimization of epidermal turnover and uniform chromophore distribution. Type 2 rejuvenation is deeper and targets decreased collagen, disorganized glycosaminoglycans and elastin, and superficial rhytides. Rejuvenation type 3 is deepest and targets deep dermal collagen disorders and skin laxity as well as loss and redistribution of subdermal fat. For each of these rejuvenation goals, various scientifically proven procedures are available. However, combining the right techniques is sometimes difficult, as improvements in established modalities and new developments are continuously evolving.

TYPE 1 REJUVENATION: EPIDERMAL TURNOVER AND UNIFORM CHROMOPHORE DISTRIBUTION

Treatment modalities for type 1 rejuvenation can be divided into superficial peeling systems and nonablative systems. Although both groups target pigment irregularities, peeling systems have a slight advantage in improving skin texture and radiance, whereas nonablative systems can additionally improve vascular conditions such as erythema or telangiectasia.

Superficial Peeling Systems

Superficial chemical peels affect the epidermis and dermal-epidermal interface; they can be used in nearly all skin types.⁵ They decrease corneocyte adhesion and can increase dermal collagen.^{6,7} Regeneration can be expected within 3 to 5 days. Common agents for superficial peels today include hydroxy acids, such as glycolic acid; salicylic acid; β -lipohydroxy acid; or tretinoin.⁸ In a concentration of 10% to 20%, trichloroacetic acid also can be used for superficial peels.

Microdermabrasion is a minimally invasive procedure that relies on an abrasive component and a vacuum component. Most systems use a handpiece embedded with a coarse inert crystal that is applied to the skin surface.⁹ Even though the ablation affects only the stratum corneum, a study by Karimipour et al¹⁰ demonstrated increased type I collagen deposition in the upper dermis after a single treatment. Clinical evidence is given for the improvement of skin texture, fine wrinkles, and dyspigmentation.^{11,12}

A new modality for superficial ablation is laser micropeeling. Based on a variable-pulse erbium:YAG (Er:YAG) laser with a wavelength of 2950 nm, tissue is vaporized

up to 50 μ m. Benefits of this modality are its tolerability under topical anesthesia, limited downtime, and high patient satisfaction with just a single procedure.¹³ Studies showed a reduction in dyschromia and low postprocedure erythema,¹⁴ but the results were not superior to intense pulsed light (IPL).¹⁵

Nonablative Systems

Short-pulsed lasers and IPL devices can be used for nonablative therapy of pigment irregularities and vascular conditions. Both modalities are based on the absorption of photons by chromophores leading to photothermolysis of the target structure. Because of the wide absorption spectrum of melanin (500–1100 nm), several laser systems with a pulse width in the nanosecond range are effective in the removal of pigmented lesions, including the pulsed dye laser (595 and 607 nm)^{16,17}; the Q-switched ruby laser (694 nm)¹⁸; the Q-switched alexandrite laser (755 nm)¹⁹; and the Q-switched Nd:YAG laser (1064 nm),²⁰ which can be frequency doubled to produce visible green light with a wavelength of 532 nm.²¹ Intense pulsed light with its wavelength spectrum of 500 to 1200 nm also targets melanin well. Because of the longer pulse durations in the millisecond range, a darkening and sloughing of the treated spots often is seen after treatments and makes IPL only second best for the treatment of pigmented lesions.²²

Oxyhemoglobin has an optimal absorption range of 577 to 660 nm and needs a higher pulse duration than melanin.²³ Good evidence has been shown for the treatment of smaller superficial vessels, often seen in telangiectasia²⁴ or poikiloderma,^{25,26} with microsecond pulsed dye lasers. For the treatment of larger deeper vessels containing more deoxyhemoglobin, longer wavelengths around 940 nm are needed. Because these vessels absorb light of the near-infrared spectrum, they are best treated with Er:YAG lasers.²⁷ Intense pulsed light with its wide spectrum (410–1400 nm) and millisecond pulse has been proven to treat small, more superficial vessels,²⁶ as well as deeper bigger vessels.²⁸

TYPE 2 REJUVENATION: DERMAL FIBER NETWORK

The second type of rejuvenation primarily targets the fiber network of the papillary dermis to improve decreased collagen as well as glycosaminoglycans and disorganized elastin. Light-based systems with wavelengths in the infrared spectrum and radiofrequency (RF) devices are particularly suitable to reach this deeper layer with only partial or without epidermal ablation, accompanied by a short downtime. Good results can be expected with 1064-nm Nd:YAG lasers using high-power micropulses,

high repetition, and large spot size to obtain bulk heating. Multiple clinical studies employing this method have demonstrated mild but reproducible improvement in rhytides with histologic evidence of neocollagenesis after treatment.^{29,30} For the treatment of photoaged skin with the fractional 1550-nm Er:YAG laser, clinical evidence is given for the treatment of wrinkles and dyschromia at the neck, décolletage,³¹ and hands (Figure 1).³² A new fractional dual-wavelength laser combining 1550-nm Er:YAG and 1927-nm thulium:YAG fibers promises even better results for the treatment of nonfacial photodamage.³³ The 1927-nm wavelength has a higher absorption coefficient for water than the 1550-nm wavelength, accompanied by a greater ability to target epidermal processes such as dyspigmentation and dyschromia.

In addition to its superficial effects on photoaged skin, IPL also affects the collagen in the upper dermis. Analysis showed that after a single IPL treatment, collagen was increased, elastin content was decreased, and elastin fiber arrangement was improved.³⁴ These histologic findings correlate with the clinical improvements seen in one other study.³⁵ The combination of IPL with RF leads to a synergistic effect, resulting in further improvement of wrinkles

and skin texture accompanied by a reduction of skin laxity.³⁶ The energy of the light source preheats the target through selective tissue absorption, resulting in lowered impedance and providing a preferential conductive pathway for RF.

TYPE 3 REJUVENATION: DEEP RHYTIDES, SKIN LAXITY, AND VOLUME LOSS

The targets of type 3 rejuvenation are missing and disorganized collagen fibers in the deep dermis as well as subcutaneous volume loss due to UV-triggered lipotrophy.

Rhytides and Skin Laxity

Several different platforms are in clinical use for the treatment of skin laxity. They can be separated into light-based systems such as lasers and broad-spectrum light sources, systems based on ultrasound or RF, and platforms combining different approaches.

Light-Based Systems—Light-based systems for the treatment of deep wrinkles and skin laxity need wavelengths that are strongly absorbed by tissue water. Particularly suitable lasers are the CO₂ (10,600 nm),



Figure 1. Photoaged skin on the left hand before (A) and after 2 treatments with the fractional 1550-nm erbium:YAG laser (B).

the 2790 nm erbium:yttrium scandium gallium garnet (Er:YSGG), or the 2940 nm Er:YAG. They can penetrate deep into the dermis independently from hemoglobin and melanin. Depending on pulse duration, fluence, and number of passes, they provide the ability to thermally ablate controlled layers of tissue with accuracy. Although CO₂ lasers are still the gold standard for skin rejuvenation, Er:YAG and Er:YSGG lasers offer a more precise ablative quality with minimal thermal damage to the surrounding tissues, resulting in less severe side effects and faster overall healing times.³⁷ Because of the prolonged recovery time and substantial risk for postoperative hyperpigmentation and hypopigmentation, scars, and other complications of ablative laser rejuvenation, fractional microablative resurfacing has become more important.

In contrast to ablative skin resurfacing in which a confluent uniform patch of dermal injury is induced, fractional resurfacing thermally ablates microscopic columns of epidermal and dermal tissue in regularly spaced arrays over a fraction of the skin surface. After a 48- to 72-hour phase of acute thermal damage, a phase of reepithelialization and repair starts, which is mediated by the adjacent columns of intact tissue. In this 30-day phase, the areas of thermally ablated tissue are repopulated by fibroblast-derived neocollagenesis and epidermal stem cell reproduction.³⁸ Because of the combination of ablation, coagulation, and deep dermal heating, the advantages of microablative fractional rejuvenation are collagen neosynthesis, tissue shrinkage, and enhanced collagen tightening.³⁹

Another light-based system for treating the photoaged body is an infrared light source with a broad spectrum of 1100 to 1800 nm. It works by heating the dermis to achieve shrinkage and neogenesis of collagen fibers. Several clinical studies shows distinct reduction of skin laxity in facial and nonfacial photoaged skin.⁴⁰⁻⁴² A histologic study by Tanaka et al⁴³ show an increase in collagen and elastin levels up to 90 days after radiation. For the treatment of the photoaged body, fluences of 37 to 47 J/cm² with 2 to 4 passes are common.

Ultrasound Systems—An alternative to light-based systems is the use of intense ultrasound. This platform works by selective thermal coagulative change within the focal region of the beam while leaving the remaining regions unaffected.⁴⁴ The heating of the tissue is caused by friction developed between the molecules due to the vibration induced by the ultrasound waves. The system creates approximately 1-mm³ zones of dermal coagulation at 3- to 4.5-mm depths depending on different probes with 4 to 7 MHz. The same transducer that delivers the focused heat to the dermis also is used to get a sonographic image of the skin to target the right depth, which is unique to

this platform. Intense ultrasound is histologically⁴⁵⁻⁴⁷ and clinically⁴⁸ proven to coagulate collagen and thereby tighten the facial and nonfacial skin. It is a safe and nonablative system with very little discomfort. Side effects are mild to moderate erythema immediately following treatment and, more rarely, elevated white linear striations.⁴⁸

Radiofrequency Devices—Radiofrequency devices are another nonablative treatment modality for UV-induced skin laxity of the body. Radiofrequency devices produce a current that is passed through the skin, thereby causing heat from tissue resistance. The heating causes direct collagen contraction and immediate skin tightening.^{49,50} Subsequent remodeling, reorientation of collagen bundles, and formation of new collagen due to wound healing is achieved over months after the treatment.^{49,51} Several technologies currently on the market vary based on the number of electrodes. Unipolar systems have 1 electrode and a grounding pad, which is placed beneath the body. As the current passes from the electrode to the grounding pad, the penetration is deep and a high temperature develops near the electrode, requiring intense epidermal cooling.⁵² In clinical studies, unipolar RF proved effective in the treatment of skin laxity and rhytides.^{53,54} Known complications are discomfort, persistent erythema, nodularity/panniculitis, burns, and dermal adhesions.

Bipolar RF systems have 2 electrodes at a fixed distance with the current passing from 1 electrode to the other. The main advantage is the controlled distribution of the current inside the tissue, as the depth of penetration is half the distance between the electrodes.⁵² Even if the depth of penetration is superficial, active cooling is necessary to prevent epidermal burns. The clinical indications and effects achieved are similar to unipolar RF⁵⁵ but with a lower risk profile and less discomfort. A recent development is a multifrequency bipolar system, which combines the frequencies 0.8, 1.7, and 2.4 MHz to target specific tissue depths. The clinical evaluation for this new platform is underway but not yet published.

Multipolar devices have 3 to 6 electrodes controlled by 1 to 3 RF generators. Although their penetration is superficial, they have deep dermal volumetric heating and no cooling is needed. Furthermore, they are less painful than unipolar or bipolar systems.⁵⁶ Their efficacy and tolerance in skin tightening and stimulation of collagen neosynthesis is shown in several studies.^{57,58} A further stage is the combination of multipolar RF and magnetic pulses. Although RF increases collagen and elastin synthesis, the magnetic pulse field should increase the production of fibroblast growth factor β 2.⁵⁹ The benefits of this platform are lower overall energy, increased safety, and no need for pretreatment preparation and topical cooling agents.

Volume Loss

After focusing on rough skin, discoloration, vascular lesions, wrinkles, and skin laxity, the last important step in 3-dimensional rejuvenation is to augment volume loss. Volume loss due to photoaging affects the hands, but the décolletage should be mentioned too, as the treatment of rhytides could improve the overall appearance. Most qualified for this approach are fillers such as hyaluronic acid and autologous fat injection treatments with poly-L-lactic acid and calcium hydroxylapatite (CaHA).

Poly-L-lactic acid is a biocompatible, biodegradable, immunologically inert, semipermanent soft tissue filler that stimulates local fibroblasts to promote neocollagenesis.⁶⁰ It is placed into the reticular dermis and subcutaneous tissue planes and lasts up to 2 years or more. Its efficacy in the treatment of nonfacial photoaging has been shown for the hands,⁶¹ neck,⁶² and chest. Adverse events for poly-L-lactic acid (eg, ecchymoses, edema, pain, pruritus, inflammation, nodules, hematomas) are not uncommon and have been reported with an incidence of 8.5% (11/130) for the face, hands, chest, or thighs.⁶³

Because synthetic CaHA is identical in chemical composition to the inorganic constituent of teeth and bone, it is nonantigenic, nonirritating, and nontoxic. The particles have a diameter of 25 to 45 μm and are blended in a gel that contains water, glycerin, and sodium carboxymethylcellulose.⁶⁴ After injection, the CaHA particles are fixed in place with thin connective tissue without reaction in surrounding tissue, evidence of migration, or evidence of heterotopic bone growth.⁶⁴ After 18 months, the microspheres begin to degrade into metabolites consisting of calcium and phosphate ions. Besides the use for facial augmentation, CaHA is particularly suitable for the treatment of volume loss in the hands, as it produces a smooth and natural-looking result (Figure 2).⁶⁵ Studies showed good results with only some temporary swelling as a side effect.^{66,67}

CONCLUSION

Three-dimensional rejuvenation of the photoaged body is an emerging area of aesthetic dermatology. The concept of specific approaches to achieve 3 different types of



Figure 2. Volume loss on the left hand before (A) and after injection of calcium hydroxylapatite (B).

rejuvenation gives a precise tool to the attending physician. Combining physical modalities with dedicated fillers to treat discolorations, elastosis, deep wrinkles, and laxity, as well as inflammation and vascular lesions, leads to enhanced clinical results and greater patient satisfaction.

REFERENCES

1. Oikarinen A. The aging of skin: chronoaging versus photoaging. *Photodermatol Photoimmunol Photomed*. 1990;7:3-4.
2. Bernstein EF, Schwartz M, Viehmeier R, et al. Measurement of protection afforded by ultraviolet-absorbing window film using an in vitro model of photodamage. *Lasers Surg Med*. 2006;38:337-342.
3. Kim EJ, Kim YK, Kim JE, et al. UV modulation of subcutaneous fat metabolism [published online ahead of print May 12, 2011]. *J Invest Dermatol*. doi:10.1038/jid.2011.106.
4. Sadick N, Marshall S, Dinkes A. *The New Natural: Your Ultimate Guide to Cutting-Edge Age Reversal*. New York, NY: Rodale Inc; 2011.
5. Rendon MI, Berson DS, Cohen JL, et al. Evidence and considerations in the application of chemical peels in skin disorders and aesthetic resurfacing. *J Clin Aesthet Dermatol*. 2010;3:32-43.
6. Van Scott EJ, Yu RJ. Hyperkeratinization, corneocyte cohesion, and alpha hydroxy acids. *J Am Acad Dermatol*. 1984;11(5, pt 1): 867-879.
7. Ditre CM, Griffin TD, Murphy GF, et al. Effects of alpha-hydroxy acids on photoaged skin: a pilot clinical, histologic, and ultrastructural study. *J Am Acad Dermatol*. 1996;34(2, pt 1):187-195.
8. Coleman WP III, Brody HJ. Advances in chemical peeling. *Dermatol Clin*. 1997;15:19-26.
9. Karimipour DJ, Karimipour G, Orringer JS. Microdermabrasion: an evidence-based review. *Plast Reconstr Surg*. 2010;125:372-377.
10. Karimipour DJ, Kang S, Johnson TM, et al. Microdermabrasion: a molecular analysis following a single treatment. *J Am Acad Dermatol*. 2005;52:215-223.
11. Coimbra M, Rohrich RJ, Chao J, et al. A prospective controlled assessment of microdermabrasion for damaged skin and fine rhytides. *Plast Reconstr Surg*. 2004;113:1438-1443; discussion 1444.
12. Shim EK, Barnette D, Hughes K, et al. Microdermabrasion: a clinical and histopathologic study. *Dermatol Surg*. 2001;27:524-530.
13. Christian MM. Microresurfacing using the variable-pulse erbium:YAG laser: a comparison of the 0.5- and 4-ms pulse durations. *Dermatol Surg*. 2003;29:605-611.
14. Somoano B, Hantash BM, Fincher EF, et al. The erbium micropeel: a prospective, randomized trial of the effects of two fluence settings on facial photoaging. *J Drugs Dermatol*. 2011;10:179-185.
15. Hantash BM, De Coninck E, Liu H, et al. Split-face comparison of the erbium micropeel with intense pulsed light. *Dermatol Surg*. 2008;34:763-772.
16. Kono T, Chan HH, Groff WF, et al. Long-pulse pulsed dye laser delivered with compression for treatment of facial lentigines. *Dermatol Surg*. 2007;33:945-950.
17. Chern PL, Domankevitz Y, Ross EV. Pulsed dye laser treatment of pigmented lesions: a randomized clinical pilot study comparison of 607- and 595-nm wavelength lasers. *Lasers Surg Med*. 2010;42:705-709.
18. Sadighha A, Saatee S, Muhagheh-Zahed G. Efficacy and adverse effects of Q-switched ruby laser on solar lentigines: a prospective study of 91 patients with Fitzpatrick skin type II, III, and IV. *Dermatol Surg*. 2008;34:1465-1468.
19. Jang KA, Chung EC, Choi JH, et al. Successful removal of freckles in Asian skin with a Q-switched alexandrite laser. *Dermatol Surg*. 2000;26:231-234.

20. Choi M, Choi JW, Lee SY, et al. Low-dose 1064-nm Q-switched Nd:YAG laser for the treatment of melasma. *J Dermatolog Treat*. 2010;21:224-228.
21. Kilmer SL, Wheeland RG, Goldberg DJ, et al. Treatment of epidermal pigmented lesions with the frequency-doubled Q-switched Nd:YAG laser. a controlled, single-impact, dose-response, multi-center trial. *Arch Dermatol*. 1994;130:1515-1519.
22. Babilas P, Schreml S, Szeimies RM, et al. Intense pulsed light (IPL): a review. *Lasers Surg Med*. 2010;42:93-104.
23. Srinivas C, Kumaresan M. Lasers for vascular lesions: standard guidelines of care. *Indian J Dermatol Venereol Leprol*. 2011;77: 349-368.
24. Jørgensen GF, Hedelund L, Haedersdal M. Long-pulsed dye laser versus intense pulsed light for photodamaged skin: a randomized split-face trial with blinded response evaluation. *Lasers Surg Med*. 2008;40:293-299.
25. Haywood RM, Monk BE. Treatment of poikiloderma of Civatte with the pulsed dye laser: a series of seven cases. *J Cutan Laser Ther*. 1999;1:45-48.
26. Rusciani A, Motta A, Fino P, et al. Treatment of poikiloderma of Civatte using intense pulsed light source: 7 years of experience. *Dermatol Surg*. 2008;34:314-319; discussion 319.
27. Civas E, Koc E, Aksoy B, et al. Clinical experience in the treatment of different vascular lesions using a neodymium-doped yttrium aluminum garnet laser. *Dermatol Surg*. 2009;35:1933-1941.
28. Raulin C, Greve B, Grema H. IPL technology: a review. *Lasers Surg Med*. 2003;32:78-87.
29. Trelles MA, Alvarez X, Martín-Vázquez MJ, et al. Assessment of the efficacy of nonablative long-pulsed 1064-nm Nd:YAG laser treatment of wrinkles compared at 2, 4, and 6 months. *Facial Plast Surg*. 2005;21:145-153.
30. Prieto VG, Diwan AH, Shea CR, et al. Effects of intense pulsed light and the 1,064 nm Nd:YAG laser on sun-damaged human skin: histologic and immunohistochemical analysis. *Dermatol Surg*. 2005;31:522-525.
31. Chiu RJ, Kridel RWH. Fractionated photothermolysis: the Fraxel 1550-nm glass fiber laser treatment. *Facial Plast Surg Clin North Am*. 2007;15:229-237, vii.
32. Sadick NS, Smoller B. A study examining the safety and efficacy of a fractional laser in the treatment of photodamage on the hands. *J Cosmet Laser Ther*. 2009;11:29-33.
33. Polder KD, Harrison A, Eubanks LE, et al. 1,927-nm fractional thulium fiber laser for the treatment of nonfacial photodamage: a pilot study. *Dermatol Surg*. 2011;37:342-348.
34. Feng Y, Zhao J, Gold MH. Skin rejuvenation in Asian skin: the analysis of clinical effects and basic mechanisms of intense pulsed light. *J Drugs Dermatol*. 2008;7:273-279.
35. Sadick NS, Weiss R, Kilmer S, et al. Photorejuvenation with intense pulsed light: results of a multi-center study. *J Drugs Dermatol*. 2004;3:41-49.
36. Sadick NS, Alexiades-Armenakas M, Bitter P Jr, et al. Enhanced full-face skin rejuvenation using synchronous intense pulsed optical and conducted bipolar radiofrequency energy (ELOS): introducing selective radiophotothermolysis. *J Drugs Dermatol*. 2005;4:181-186.
37. Alexiades-Armenakas MR, Dover JS, Arndt KA. The spectrum of laser skin resurfacing: nonablative, fractional, and ablative laser resurfacing. *J Am Acad Dermatol*. 2008;58:719-737; quiz 738-740.
38. Laubach HJ, Tannous Z, Anderson RR, et al. Skin responses to fractional photothermolysis. *Lasers Surg Med*. 2006;38:142-149.
39. Gotkin RH, Sarnoff DS, Cannarozzo G, et al. Ablative skin resurfacing with a novel microablative CO₂ laser. *J Drugs Dermatol*. 2009;8:138-144.

40. Alexiades-Armenakas M. Assessment of the mobile delivery of infrared light (1100-1800 nm) for the treatment of facial and neck skin laxity. *J Drugs Dermatol*. 2009;8:221-226.
41. Carniol PJ, Dzopa N, Fernandes N, et al. Facial skin tightening with an 1100-1800 nm infrared device. *J Cosmet Laser Ther*. 2008;10:67-71.
42. Goldberg DJ, Hussain M, Fazeli A, et al. Treatment of skin laxity of the lower face and neck in older individuals with a broad-spectrum infrared light device. *J Cosmet Laser Ther*. 2007;9:35-40.
43. Tanaka Y, Matsuo K, Yuzuriha S. Long-term evaluation of collagen and elastin following infrared (1100 to 1800 nm) irradiation. *J Drugs Dermatol*. 2009;8:708-712.
44. Kennedy JE, Ter Haar GR, Cranston D. High intensity focused ultrasound: surgery of the future? *Br J Radiol*. 2003;76:590-599.
45. Gliklich RE, White WM, Slayton MH, et al. Clinical pilot study of intense ultrasound therapy to deep dermal facial skin and subcutaneous tissues. *Arch Facial Plast Surg*. 2007;9:88-95.
46. Laubach HJ, Makin IRS, Barthe PG, et al. Intense focused ultrasound: evaluation of a new treatment modality for precise microcoagulation within the skin. *Dermatol Surg*. 2008;34:727-734.
47. White WM, Makin IRS, Barthe PG, et al. Selective creation of thermal injury zones in the superficial musculoaponeurotic system using intense ultrasound therapy: a new target for noninvasive facial rejuvenation. *Arch Facial Plast Surg*. 2007;9:22-29.
48. Alam M, White LE, Martin N, et al. Ultrasound tightening of facial and neck skin: a rater-blinded prospective cohort study. *J Am Acad Dermatol*. 2010;62:262-269.
49. Zelickson BD, Kist D, Bernstein E, et al. Histological and ultrastructural evaluation of the effects of a radiofrequency-based nonablative dermal remodeling device: a pilot study. *Arch Dermatol*. 2004;140:204-209.
50. Kist D, Burns AJ, Sanner R, et al. Ultrastructural evaluation of multiple pass low energy versus single pass high energy radio-frequency treatment. *Lasers Surg Med*. 2006;38:150-154.
51. Bogle MA, Ubelhoer N, Weiss RA, et al. Evaluation of the multiple pass, low fluence algorithm for radiofrequency tightening of the lower face. *Lasers Surg Med*. 2007;39:210-217.
52. Elsaie ML, Choudhary S, Leiva A, et al. Nonablative radiofrequency for skin rejuvenation. *Dermatol Surg*. 2010;36:577-589.
53. Emilia del Pino M, Rosado RH, Azuela A, et al. Effect of controlled volumetric tissue heating with radiofrequency on cellulite and the subcutaneous tissue of the buttocks and thighs. *J Drugs Dermatol*. 2006;5:714-722.
54. el-Domyati M, el-Ammawi TS, Medhat W, et al. Radiofrequency facial rejuvenation: evidence-based effect. *J Am Acad Dermatol*. 2011;64:524-535.
55. Alexiades-Armenakas M, Dover JS, Arndt KA. Unipolar versus bipolar radiofrequency treatment of rhytides and laxity using a mobile painless delivery method. *Lasers Surg Med*. 2008;40:446-453.
56. Royo de la Torre J, Moreno-Moraga J, Muñoz E, et al. Multisource, phase-controlled radiofrequency for treatment of skin laxity: correlation between clinical and in-vivo confocal microscopy results and real-time thermal changes. *J Clin Aesthet Dermatol*. 2011;4:28-35.
57. Boisnic S, Branchet MC. Ex vivo human skin evaluation of localized fat reduction and anti-aging effect by TriPollar radio frequency treatments. *J Cosmet Laser Ther*. 2010;12:25-31.
58. Kaplan H, Gat A. Clinical and histopathological results following TriPollar radiofrequency skin treatments. *J Cosmet Laser Ther*. 2009;11:78-84.
59. Tepper OM, Callaghan MJ, Chang EI, et al. Electromagnetic fields increase in vitro and in vivo angiogenesis through endothelial release of FGF-2. *FASEB J*. 2004;18:1231-1233.
60. Gogolewski S, Jovanovic M, Perren SM, et al. Tissue response and in vivo degradation of selected polyhydroxyacids: polylactides (PLA), poly(3-hydroxybutyrate) (PHB), and poly(3-hydroxybutyrate-co-3-hydroxyvalerate) (PHB/VA). *J Biomed Mater Res*. 1993;27:1135-1148.
61. Palm MD, Woodhall KE, Butterwick KJ, et al. Cosmetic use of poly-L-lactic acid: a retrospective study of 130 patients. *Dermatol Surg*. 2010;36:161-170.
62. Mazzucco R, Hexsel D. Poly-L-lactic acid for neck and chest rejuvenation. *Dermatol Surg*. 2009;35:1228-1237.
63. Peterson JD, Goldman MP. Rejuvenation of the aging chest: a review and our experience. *Dermatol Surg*. 2011;37:555-571.
64. Tzikas TL. Evaluation of the Radiance FN soft tissue filler for facial soft tissue augmentation. *Arch Facial Plast Surg*. 2004;6:234-239.
65. Marmur ES, Phelps R, Goldberg DJ. Clinical, histologic and electron microscopic findings after injection of a calcium hydroxylapatite filler. *J Cosmet Laser Ther*. 2004;6:223-226.
66. Busso M, Applebaum D. Hand augmentation with Radiesse (calcium hydroxylapatite). *Dermatol Ther*. 2007;20:385-387.
67. Gargasz SS, Carbone MC. Hand rejuvenation using Radiesse. *Plast Reconstr Surg*. 2010;125:259e-260e. ■