A SUPPLEMENT TO

DERNATOLICE DERNATOLICE

February 2008 Volume 21 No. 2 S1



Dermal Fillers for Facial Rejuvenation and Restoration: Integrating New **Therapies Into Clinical Practice**



This activity is jointly sponsored by Postgraduate Institute for Medicine and The MedCom Resource, Inc.

> Supported by an educational grant from Dermik Laboratories, a business of sanofi-aventis U.S. LLC.

> > Designated for 1.25 AMA PRA Category 1 Credits™

Editor

Caroline Colyer 973-206-8095/fax 973-206-9256 E-mail: caroline.colyer@qhc.com

Associate Editor

Jenn A. Verlangieri 973-206-8094/fax 973-206-9256 E-mail: jennifer.verlangieri@qhc.com

Proofreader Michele V. Murray

Wileliese V. Walla

Creative Director Mary Ellen Niatas 973-206-8973

Production Manager Jaime Serra 973-206-8011

Subscriber Inquiry Line 800-976-4040

Vice President/Group Publisher Sharon Finch 973-206-8952

Publishing Consultant Claudia Shayne-Ferguson

Program Manager Shannon C. Conover 973-206-8015

Classified Advertising Valley Forge Publishing

866-312-8805

Reprint Inquiries Shannon C. Conover 973-206-8015 E-mail: shannon.conover@qhc.com

Director, Marketing Research Lori Raskin

QUADRANT HEALTHCOM INC. President & CEO Stephen Stoneburn



7 Century Dr Suite 302 Parsippany, NJ 07054-4609 A Supplement to

COSMETIC DERMATOLOGY®

February 2008 Volume 21 No. 2 S1

- 3 Combining Advanced Injection Techniques: Integrating New Therapies Into Clinical Practice Wm. Philip Werschler, MD
- 7 Current Treatment Options: What's In and What's Out? Gary D.Monheit, MD
- 11 Choosing the Most Appropriate Filler: Safety, Techniques, and Combinations Chérie M. Ditre, MD
- 15 CME Test
- 16 Application for Category I Credits

PUBLISHER: Cosmetic Dermatology® (ISSN 1041-3766) (GST #128741063) is published monthly by Quadrant HealthCom Inc., with editorial and advertising offices at 7 Century Dr, Suite 302, Parsippany, NJ 07054-4609; fax 973-206-9378. Periodicals postage paid at Parsippany, NJ, and additional mailing offices.

 $Copyright @2008. Quadrant \ Health Com \ Inc. \ All \ rights \ reserved.$

Dermal Fillers for Facial Rejuvenation and Restoration: Integrating New Therapies Into Clinical Practice

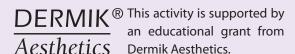
Release date: February 2008 Expiration date: February 2009

Estimated time to

complete activity: 1 hour 15 minutes



Jointly sponsored by Postgraduate Institute for Medicine and The MedCom Resource, Inc.



TARGET AUDIENCE

This activity has been designed to meet the educational needs of cosmetic dermatologists and aesthetic surgeons involved in the management of patients with facial aging.

STATEMENT OF NEED

Facial lipoatrophy is a natural part of the aging process. It is important to address this condition, as it can stigmatize patients, causing them to suffer lower self-esteem and quality of life. As newer products become available, there is an ongoing need for physician education. This continuing medical education (CME) program will assist physicians in understanding how to combine toxins and fillers as part of an overall plan to achieve facial rejuvenation and restoration of patients in their practices. The authors discuss the treatments available to meet this goal.

EDUCATIONAL OBJECTIVES

After completing this activity, the participant should be better able to:

- 1. Define the techniques of assessment for the aging face
- 2. Identify a personal plan for diagnosing and prescribing therapy for the aging face

- 3. Describe the use of collagen stimulators for nonsurgical treatment of facial rejuvenation and restoration
- 4. Identify treatment options based on location and stage of lipoatrophy
- 5. Describe safety considerations and injection techniques for dermal fillers

FACULTY

Chérie M. Ditre, MD, is Director, Cosmetic Dermatology & Skin Enhancement Center, Penn Medicine at Radnor, Department of Dermatology, Pennsylvania. Gary D. Monheit, MD, is President, Total Skin and Beauty Dermatology Center, and Associate Clinical Professor, Departments of Dermatology and Ophthalmology, University of Alabama at Birmingham. Wm. Philip Werschler, MD, is Assistant Clinical Professor of Medicine/Dermatology, University of Washington School of Medicine, Seattle.

ACCREDITATION STATEMENT

This activity has been planned and implemented in accordance with the Essential Areas and Policies of the Accreditation Council for Continuing Medical Education (ACCME) through the joint sponsorship of Postgraduate Institute for Medicine (PIM) and The MedCom Resource, Inc. PIM is accredited by the ACCME to provide CME for physicians.

CREDIT DESIGNATION

PIM designates this educational activity for a maximum of 1.25 AMA PRA Category 1 CreditsTM. Physicians should only claim credit commensurate with the extent of their participation in the activity.

DISCLOSURE OF CONFLICTS OF INTEREST

PIM assesses conflict of interest with its instructors, planners, managers and other individuals who are in a position to control the content of

CME INFORMATION

CME activities. All relevant conflicts of interest that are identified are thoroughly vetted by PIM for fair balance, scientific objectivity of studies utilized in this activity, and patient care recommendations. PIM is committed to providing its learners with high-quality CME activities and related materials that promote improvements or quality in health care and not a specific proprietary business interest of a commercial interest.

The faculty reported the following financial relationships or relationships to products or devices they or their spouse/life partner have with commercial interests related to the content of this CME activity:

Dr. Ditre is a speaker for Dermik Aesthetics. Dr. Monheit is a consultant for Allergan, Inc; Genzyme Corporation; and Medicis Pharmaceutical Corporation; and a researcher for Allergan, Inc; ColBar LifeScience Ltd; and Medicis Pharmaceutical Corporation. Dr. Werschler is a consultant for Allergan, Inc; BioForm Medical, Inc; Dermik Laboratories; Medicis Pharmaceutical Corporation; MyoScience Inc; and OrthoNeutrogena; a researcher for Allergan, Inc; BioForm Medical, Inc; Dermik Laboratories; and Medicis Pharmaceutical Corporation; and a stockholder of Allergan, Inc; BioForm Medical, Inc; and MyoScience Inc.

The planners and managers reported the following financial relationships or relationships to products or devices they or their spouse/life partner have with commercial interests related to the content of this CME activity:

The following planners and managers, Jaime Pedro; Liza Risoli; and John Russo Jr, PharmD, have no real or apparent conflicts of interest to report.

The following PIM clinical content reviewers, Jan Hixon, RN; Trace Hutchison, PharmD; and Linda Graham, RN, have no real or apparent conflicts of interest to report.

METHOD OF PARTICIPATION

There are no fees for participating in and receiving CME credit for this activity. During the period February 2008 through February 2009, participants must 1) read the learning objectives and

faculty disclosures; 2) study the educational activity; 3) complete the posttest by recording the best answer to each question in the answer key on the evaluation form; 4) complete the evaluation form; and 5) mail or fax the evaluation form with answer key to PIM.

A statement of credit will be issued only upon receipt of a completed activity evaluation form and a completed posttest with a score of 70% or better. Your statement of credit will be mailed to you within 3 weeks.

MEDIA

Printed supplement

DISCLOSURE OF UNLABELED USE

This educational activity may contain discussion of published and/or investigational uses of agents that are not indicated by the US Food and Drug Administration. PIM, The MedCom Resource, Inc, and Dermik Aesthetics do not recommend the use of any agent outside of the labeled indications.

The opinions expressed in the educational activity are those of the faculty and do not necessarily represent the views of PIM, The MedCom Resource, Inc, and Dermik Aesthetics. Please refer to the official prescribing information for each product for discussion of approved indications, contraindications, and warnings.

DISCLAIMER

Participants have an implied responsibility to use the newly acquired information to enhance patient outcomes and their own professional development. The information presented in this activity is not meant to serve as a guideline for patient management. Any procedures, medications, or other courses of diagnosis or treatment discussed or suggested in this activity should not be used by clinicians without evaluation of their patients' conditions and possible contraindications on dangers in use, review of any applicable manufacturer's product information, and comparison with recommendations of other authorities.

Combining Advanced Injection Techniques: Integrating New Therapies Into Clinical Practice

Wm. Philip Werschler, MD

Durability and longevity of effect are significant considerations in the choice of treatment for cosmetic dermatology patients. Equally important, if not more so, is the utility of a product to achieve and maintain an effect. In this supplement, my colleagues Gary Monheit, MD, and Chérie Ditre, MD, review the current treatment choices, filler techniques, and combinations that are available. In this article, I would like to introduce the concept of regional aesthetic volume enhancement as part of a comprehensive approach for developing treatment protocols for the youthful enhancement patient seeking obvious improvement of natural features, the transitional rejuvenation patient attempting to recapture youth, and the fully adult patient seeking restoration of both health- and ageappropriate facial beauty.

oday, there are many fillers available in the United States to enhance natural features, rejuvenate fading youth, and restore aged facial features (Table 1). Equally important, if not more so, in clinical practice is the ease of use and utility of a product and achieving and maintaining a desired effect over time. Through both a thorough understanding of the mechanism of action of each product category and a judicious application of their unique features and benefits, it is possible to meet reasonable treatment expectations for

Presented in part at the sixth annual meeting of the American Society of Cosmetic Dermatology & Aesthetic Surgery, Las Vegas, Nevada, December 1, 2007.

the vast majority of patients seeking this nonsurgical total facial rejuvenation and restoration approach to aesthetic treatment.

REGIONAL AESTHETIC VOLUME ENHANCEMENT

Regional aesthetic volume enhancement (RAVE) compartmentalizes facial areas or regions as discrete cosmetic units known as facial treatment zones (Figure 1). This approach facilitates a conceptual predetermination of the outcome desired for volume enhancement and aids in the selection, placement, and injection techniques of differing facial-shaping agents.

Inherent to this approach is the diagnosis of the relative contributory components of the aging face, with a treatment plan of which products are to be used in which areas and for what effect prior to their prescribing. This process requires effective communication between the provider and patient prior to any actual treatment, in a process best described by Werschler and Fried³ in the CPS-D/STEP (cosmetic procedure screen—dermatology/stress, target, envision, proactive) protocol.

RAVE is best considered in the context of a comprehensive approach to desired cosmetic outcomes, whether they are principally for enhancement, rejunvenative, or restorative purposes.

Whereas the current overwhelming consumer trend is to favor nonsurgical procedures over surgical

TABLE 1

Application of Facial-Shaping Agents

- Enhance natural features
- · Rejuvenate fading youth
- Restore aged facial features

Figure not available online

Figure 1. Regional aesthetic volume enhancement compartmentalizes facial areas or regions as discrete cosmetic units, or facial treatment zones, and seeks to rebalance and reharmonize facial features both within and between zones to achieve the desired cosmetic outcome. Illustration courtesy of Irene Matiatos Russo, PhD.

procedures, RAVE and nonsurgical total facial rejuvenation and restoration as volumetric-lifting procedures are alternatives and complement traditional surgical tension-lifting procedures (Table 2).

Through careful examination, assessment, and analysis of the patient's underlying contributory factors (eg, biometric volume loss, the aging dermal effects of photodamage), RAVE can assist in restoring the loss of youthful balance, symmetry, and harmony leading to geometric inversion of the facial triangle in order to achieve desired contours and minimize the atrophy and redistribution of subsurface tissues that combine to result in an aged appearance (Table 3). The goal

TABLE 2

Regional Aesthetic Volume Enhancement

- A technique concept, not product dependant
- A method of assessing, planning, and performing volume enhancement to replace (augment) biometric volume loss/alteration—a primary intrinsic component of the aging face
- Uses facial-shaping agents to reharmonize, rebalance and restore symmetry to the facial treatment zones
- The volume component volumetric lifting of nonsurgical total facial rejuvenation and restoration

with RAVE is restoration of facial volume and contours rather than simply filling in wrinkles and lines.

RAVE IN RELATION TO PATIENT AGE

The issues faced by cosmetic dermatology patients follow a spectrum of fairly simple and straightforward complaints from aging to more complex issues. Although facial changes occur gradually as a continuum, in practice there are 3 fairly disparate groups that can be defined psychodynamically by their respective motivational drivers: the youthful enhancement patient, the transitional rejuvenation patient, and the fully adult restoration patient (Figure 2). It is instructive to understand the needs and expectations of each group.

Youthful Enhancement Patient

The youthful enhancement patient population generally ranges from 18 to 35 years of age. They seek enhancement of certain features with a goal of maximizing natural youth and beauty. For example, the patients might consider the lips too small, or the mental crease too distinctive. Also, the nasal tip might be a bit asymmetric, the eyebrows flatter than desired, or the cheeks somewhat depressed. The youthful enhancement patient may also request the addition of features that he or she never had, such as enhanced cheek projection, the appearance of wider eyes, or voluptuous lips.

Success is achieved when the end result is aesthetic improvement (or enhancement) from the youthful patient's perspective—creating natural-looking contours or augmenting more modest physical traits. This nonsurgical option has advantages compared to permanent implants so that as fashions or desires change, the effect of the enhancement gradually resolves if not maintained. Nonpermanent options also allow for subsequent modifications to meet age- and appearance-related changes that will occur

TABLE 3

Cumulative Changes in Facial Geometry Over Time

- Dermal atrophy
- Muscle atrophy
- Lipoatrophy and redistribution (ie, lipoalteration)
- Bone and cartilage loss and remodeling (ie, osteoalteration)
- Hairline changes

Figure not available online

Figure 2. Although facial changes occur gradually as a continuum, in practice there are 3 fairly disparate groups: the youthful enhancement patient (A), the transitional rejuvenation patient (B, C), and the fully adult restoration patient (D, E). Illustrations courtesy of Irene Matiatos Russo, PhD.

over time as a consequence of both intrinsic and extrinsic aging.

It has been said that the youthful enhancement patient seeks to look better than his or her natural appearance.

Transitional Rejuvenation Patient

Transitional rejuvenation patients are approximately 35 to 55 years of age. This age group is sometimes referred to as the "youth corridor" where, given good genetics, good health, good skin care, and good luck, there are minimal aging changes that occur in facial appearance. This group psychodynamically desires rejuvenation of features in order to recapture as much of their youthful appearance as possible. The most common structural facial changes include an overall flattening and laxity of the mid face with a more pronounced vertical dimension of the lower eyelid, overly defined nasolabial folds, and a diminution of the projection of the perioral region, including the lips. In addition, there may be pronounced marionette lines, drooping oral

commissures, and development of the prejowl sulcus with loss of the curvilinear sweep of the mandible.

Rejuvenation of these features in the mid to lower face can be accomplished by using soft tissue fillers. When dermal atrophy and laxity are evident, stimulatory fillers used for pan-facial dermal augmentation are especially helpful to repair the foundation prior to more defining contour treatments. Success is achieved when patients recapture a healthier age-appropriate appearance that is more consistent psychologically with their perceived age and youthfulness and physiologically with their physical conditioning.

It is has been said of this group that the desire is to look as good as they feel.

Fully Adult Restoration Patient

Patients who are 55 years of age or older comprise this final demographic. They experience marked changes in the upper, mid, and lower face. The original triangular facial shape has morphed to a trapezoidal shape or perhaps even to an inverted triangle.

Correction of descent of the facial soft tissues will help return the patient to a more aesthetically appealing, age-appropriate appearance. However, these patients may also require treatment to return the skin to a healthy appearance as well. This is especially true following disease-related events such as skin cancer, precancerous lesions, and rosacea.

Typically, these patients have experienced significant amounts of both intrinsic and extrinsic (photodamage) aging changes and will commonly need dermal structural support and volume replacement with collagen stimulating fillers as a baseline prior to contouring specific features. Additionally, resurfacing options to both repair and enhance surface texture and pigmentation changes resulting from age and to stimulate epidermal cell turnover are commonly employed.

Thus, multiple regions of the face, both inside and outside, must be addressed to achieve the desired outcome. In addition to the lower third of the face, especially the perioral area, the mid face is likely to require added volume. The use of facial-shaping agents in the malar and infraorbital areas, as well as the prejowl sulcus, can bring the patient closer to restoration of the triangle of beauty associated with a younger and healthier face. Typically, this population is more likely than the others to require combined surgical and nonsurgical approaches to achieve the desired end result.

Fully adult restoration patients are typified by not wanting to look especially younger, but rather especially good for their age when compared to their peers.

TREATMENT OPTIONS

Robert Jackson, MD, past president of the American Academy of Cosmetic Surgery, has stated, "The science and technology we have today allow us to do many things and create aesthetically pleasing changes for our patients."

As the resources, capabilities, and skill sets of aesthetic dermatologists continue to develop and improve, it is time to move on from simply correcting

superficial facial lines and wrinkles to a more global understanding and approach of the dynamics of facial aging. Combined with a deeper appreciation of the psychological motivations of patients seeking cosmetic procedures, a comprehensive approach to patient satisfaction can be crafted. This approach should be well within the treatment domain of aesthetic dermatologists as well as cosmetic and plastic surgeons.

The framework of nonsurgical total facial rejuvenation and restoration, using facial-shaping agents to correct and balance facial treatment zones and working in the context of RAVE where applicable, helps physicians to dynamically balance the face to address patient desires across the age spectrum. By using multiple products, each with unique attributes, in combination to create natural-looking youthful contours, symmetry, and balance, RAVE offers a more aesthetically holistic approach to optimizing facial shaping with minimally invasive approaches. RAVE is independent of any single product or injection style. Rather, it is the logical culmination of careful patient assessment, visualization of the desired outcome, and a specifically chosen armamentarium of products. In this context, fillers used alone and in combination become a foundation treatment that is complementary to toxins, lasers, and other procedures.

As part of a comprehensive approach to the cosmetic patient, careful discussion and determination of the desired outcome is essential for success. Guidelines for cosmetic dermatology patient screening are especially helpful to prevent unintended expectations and performance mismatches and to assist in keeping both the patient and provider on the same page.

REFERENCES

- Alam M, Yoo SS. Technique for calcium hydroxylapatite injection for correction of nasolabial fold depressions. J Am Acad Dermatol. 2007;56:285-289.
- 2. Felderman I.I. Radiesse[™] for facial rejuvenation. *Cosmet Dermatol.* 2005;18:823-826.
- 3. Werschler WP, Fried R. The key to mastering cosmetic dermatology patient selection. Skin & Aging. 2006;14:42-50. ■

Current Treatment Options: What's In and What's Out?

Gary D. Monheit, MD

Fillers are important tools in combating the aging phenomenon. A variety of formulations are available, and each has its benefits and limitations. In this article, popular forms of temporary synthetic injectable fillers are compared, including human collagen, hyaluronic acid, poly-L-lactic acid, and calcium hydroxylapatite. In addition, 2 nondegradable injectable fillers, silicone and polymethylmethacrylate, are compared.

uccess in nonsurgical facial rejuvenation starts by selecting the right fillers. Ultimately, success lies in recognizing the facial defects and applying expertise in injection techniques. In addition, it is essential to understand the goals of treatment from the patient's perspective. The focus of this article is on fillers available to clinicians practicing in the United States.

THE AGING PROCESS

As we age, 2 distinct types of changes combine to cause facial aging: intrinsic changes, mediated to a significant extent by genetic factors, and extrinsic changes, the results of environmental factors (eg, exposure to the sun's rays).

Intrinsic aging is a continuous process whereby skin and collagen production slows and changes in elastin result in a loss of tone. Dead skin cells do not shed as quickly, and the turnover of new skin cells may decrease slightly. Although these changes begin during the third decade of life, the signs of intrinsic aging are usually

Presented in part at the sixth annual meeting of the American Society of Cosmetic Dermatology & Aesthetic Surgery, Las Vegas, Nevada, December 1, 2007.

not visible for many years thereafter. By comparison, extrinsic aging is caused by sun exposure (photoaging), with contributions by repetitive facial expressions, gravity, sleeping positions, and smoking (Table 1).¹

The extent of photoaging depends on a person's skin color and history of sun exposure over many years. Fair-skinned individuals with a history of sun exposure are at greatest risk for developing the signs of photoaging, which include fine wrinkles and a mottled complexion.

Repeated UV exposure breaks down collagen and impairs collagen synthesis. Changes in elastin (as in intrinsic aging) result in a loss of tone. In addition, the skin becomes loose, wrinkled, and leathery much earlier with unprotected exposure to sunlight.¹

CHOOSING THE PROPER FILLER

A variety of filler formulations are available for managing the aging phenomenon. Each has its benefits and limitations. Table 2 lists the fillers that have been introduced in the United States over the past 35 years.²

HUMAN-DERIVED COLLAGEN

Human-derived collagens (ie, CosmoDerm and CosmoPlast) are harvested from bioengineered human skin cells and then seeded on a 3-dimensional mesh that is identical to the human dermis but lacks immune cells.³

CosmoDerm is injected into the upper papillary dermis to correct fine lines. CosmoPlast is used to treat deeper rhytides, smooth scars, and enhance the lips. Unlike CosmoDerm, CosmoPlast is cross-linked with glutaraldehyde to increase its strength and longevity.³ The results are immediate and last 2 to 5 months.⁴ Initial swelling resolves in a few days and can be minimized by avoiding strenuous exercise, alcoholic beverages, and sun exposure in the first 24 hours.

Neither product requires skin testing prior to use. However, patients with an allergy to lidocaine or with a history of anaphylaxis to any allergen should not receive CosmoDerm or CosmoPlast. These products should also be used cautiously in patients with

TABLE 1

Facial Signs of Intrinsic and Extrinsic Aging¹

Type of Aging	Signs	
Intrinsic	Fine wrinkles Thin, transparent skin Loss of underlying fat Hollowed cheeks and eye sockets Loss of firmness on the hands and neck Bone loss Sagging skin Dry skin	
Extrinsic	Dry texture Fine and coarse wrinkling Sallow color, dyspigmentation Loss of tone	

autoimmune diseases such as lupus, scleroderma, or rheumatoid arthritis.³ A summary of the uses of human collagen and common adverse reactions are listed in Table 3.

TEMPORARY SYNTHETIC INJECTABLE FILLERS

Three popular forms of temporary synthetic injectable fillers are hyaluronic acid, poly-L-lactic acid, and calcium hydroxylapatite.

Upon injection, hyaluronic acid fillers provide an immediate response. However, the duration of effect is the shortest among the temporary synthetic injectable fillers—between 3 and 9 months.⁶

Hyaluronic acid is extremely hydrophilic, binding water up to 1000 times its volume and attracting water into the extracellular matrix that produces turgor. There are 4 available forms of hyaluronic acid injectables in the United States that have been approved by the US Food and Drug Administration: Captique, Hylaform, Juvéderm, and Restylane. Table 4 lists their uses and adverse reactions.

Despite the fact that each product is hyaluronic acid, they are not identical formulations. However, each may be used to fill the nasolabial folds, glabellar lines, and distensible scars. They are not recommended for use on the lips. Although rare, there have been reports of delayed inflammatory skin

TABLE 2

Introduction of Fillers in the United States since 1972

Year	Product
1972–1975	Collagen Fibril
2002	CosmoDerm, CosmoPlast (collagen) Restylane, Hylaform (hyaluronic acid)
2004	Calcium hydroxylapatite Poly-L-lactic acid
2006	Juvéderm (hyaluronic acid) Polymethylmethacrylate microspheres

reactions following hyaluronic acid filler procedures. Other adverse reactions include transient erythema, localized swelling, bruising, and tenderness. If the hyaluronic acid is injected too superficially in the dermis, there may be an apparent blue mark caused by the Tyndall effect.³

Two products—poly-L-lactic acid and calcium hydroxylapatite—are longer lasting than collagen (Table 5). They are sometimes referred to as semipermanent fillers, reflecting their comparatively durable response.^{8,9}

TABLE 3

Uses of Human Collagen and Common Adverse Events⁵

Clinical Use

Rhytides, fine and coarse Atrophic scars Lip augmentation Folds, grooves

Adverse Reaction

Soft tissue swelling Bruising Skin necrosis (rare) Blindness (remote)

TABLE 4

Uses of Short-Acting Hyaluronic Acid Fillers and Common Adverse Events⁵

Clinical Use

Coarse rhytides
Atrophic scars
Lip augmentation
Folds and grooves

Adverse Reaction

Soft tissue swelling Bruising Postoperative discomfort (rare)

Volume restoration with poly-L-lactic acid is gradual and incremental with subsequent treatments. Generally, 3 to 6 sessions are needed, and the results last up to 2 years with repeated treatment. The most commonly observed adverse event associated with the use of poly-L-lactic acid is delayed occurrence of subcutaneous papules at the injection site, which are typically palpable, asymptomatic, and nonvisible.^{5,9}

By comparison, the response to calcium hydroxylapatite is related to the volume injected. Clinical results last for 9 to 12 months. Calcium hydroxylapatite is generally well tolerated. However,

precise technique is required with lip augmentation in order to reduce the risk of submucosal nodules, which occur in approximately 10% of patients treated in the lips. Most often the nodules are not visible and resolve without intervention ¹⁰

NONDEGRADABLE INJECTABLE FILLERS

Silicone and polymethylmethacrylate are synthetic products that are not readily metabolized and removed by the body.^{11,12}

Silicone is used for the treatment of atrophic scars, as well as lip and nasolabial fold augmentation. Polymethylmethacrylate is a permanent injectable implant for the treatment of soft tissue defects of the face. The value of a permanent product is obvious. However, facial changes that occur with aging continue after the nondegradable fillers are in place. Therefore, additional treatment might be required over time, and removal of these products is accomplished with difficulty, if at all.

CONCLUSIONS

Many treatment options are available for nonsurgical facial rejuvenation. With careful assessment, the appropriate intervention can be identified and employed to correct the inevitable effects of aging on appearance.

REFERENCES

- AgingSkinNet. Causes of aging skin. http://www .skincarephysicians.com/agingskinnet/basicfacts.html. Accessed December 18, 2007.
- 2. Haneke E. Skin rejuvenation without a scalpel, I: fillers. *J Cosmet Dermatol.* 2006;5:157-167.
- 3. Fernandez EM, Mackley CL. Soft tissue augmentation: a review. *J Drugs Dermatol.* 2006;5:630-641.

	Λ.	ы		
	Λ.	124		•

Longer-Acting Injectable Fillers⁵

Compound	Onset	Duration	Adverse Reactions
Calcium hydroxylapatite	Immediate	15–18 mo	Bruising Erythema Nodule formation (especially in lips) Granulomas If injected on bone (under periosteum), bone formation can occur
Poly-L-lactic acid	Full effects in weeks to months	Up to 24 mo with repeat treatments	Bruising Erythema Nodule formation Granulomas

TREATMENT OPTIONS

- 4. Lowe N. New filler agents: what can we learn from Europe? *Pract Dermatol.* 2004;1:29-33.
- Sengelman RD. Exploring management options for facial lipoatrophy: focus on semipermanent fillers. March 26, 2006. http://www.medscape.com/viewarticle/527948_12. Accessed December 18, 2007.
- Werschler WP, Weinkle S. Longevity of effects of injectable products for soft-tissue augmentation. J Drugs Dermatol. 2005;4:20-27.
- 7. Haake EM, Holbrook K. The structure and development of the skin. In: Freedberg I, Fitzpatrick TB, eds. Fitzpatrick's
- Dermatology in General Medicine. 5th ed. New York, NY: McGraw-Hill; 1999:70-86.
- 8. Radiesse [package insert]. San Mateo, CA: BioForm Medical; 2006.
- 9. Sculptra [package insert]. Bridgewater, NJ: sanofi-aventis; 2004.
- Tzikas TL. Evaluation of the Radiance FN soft tissue filler for facial soft tissue augmentation. Arch Facial Plast Surg. 2004;6:234-239.
- 11. Silikon 1000 [package insert]. Fort Worth, TX: Alcon Laboratories.
- 12. Artes Medical Web site. http://www.artesmedical.com/about.php. Accessed February 8, 2006.

Choosing the Most Appropriate Filler: Safety, Techniques, and Combinations

Chérie M. Ditre, MD

Offering patients only one or two treatment options for rejuvenation of the aging face is rapidly becoming insufficient. This article presents my algorithm for combining nonsurgical treatments to correct changes to the mid face and lips that accompany advanced aged.

he most recent findings from the American Society of Plastic Surgeons¹ and the American Academy of Cosmetic Surgery² support the need for greater information and guidance on the safe and effective use of combined nonsurgical treatments for facial rejuvenation.

In this article, these relevant statistics are summarized and a personal algorithm for complete nonsurgical management of the aging face is provided as a guide to help clinicians advance their practices in this field.

INCREASE IN NONSURGICAL TREATMENTS

Data from the American Society of Plastic Surgeons survey reveal that from 2005 to 2006, the use of all fillers increased by 41% (Table 1). The comparative increase in the use of calcium hydroxylapatite and poly-L-lactic acid was approximately the same at 19% and 18%, respectively. However, the greatest increase (59%) occurred with the shorter-acting hyaluronic acid products. 1

Today, half of the members of the American Academy of Cosmetic Surgery (AACS) offer semipermanent fillers, such as poly-L-lactic acid and calcium hydroxylapatite, and more than 9 of 10 AACS members

Presented in part at the sixth annual meeting of the American Society of Cosmetic Dermatology & Aesthetic Surgery, Las Vegas, Nevada, December 1, 2007.

offer botulinum toxin and hyaluronic acid (Table 2).² When compared to cosmetic surgery procedures offered by AACS members, the semipermanent fillers are offered more often than almost all 23 surgical procedures (Table 3).

A PERSONAL ALGORITHM FOR NONSURGICAL FACIAL REJUVENATION

Offering patients only one or two filler options falls short of providing optimal management of the changes that accompany the aging face. Clinicians must identify the products that perform best for them and their patients. The following observations are based on my experience as Director of the Cosmetic Dermatology & Skin Enhancement Center at Penn Medicine at Radnor in Pennsylvania.

MIDFACE CORRECTION

The aging patient typically exhibits symmetrical mid- to lower-face depression, most notably with sunken cheeks. There is a loss of the youthful lifting projection and roundness of the zygoma with submalar depression. Treatment objectives include volumizing the cheeks, lifting of the nasolabial and mesolabial folds, and reversing the deepening of the nasolabial and nasojugal (tear trough) creases.

Volumizing the Cheeks

Poly-L-lactic acid is used as a nonsurgical cheek implant. It is important that this product be injected in the deep dermis or subcutaneous layer and that superficial injections are avoided.³ Volume restoration occurs over time, with subsequent treatments providing gradual and incremental volume to the injected area. Typically, 3 to 6 sessions are required in order to produce results that will last up to 2 years. Poly-L-lactic acid comes as a freezedried powder with 2 vials per box.³ I reconstitute the product using 5 mL sterile water for injection plus 1 mL lidocaine. During each session, typical treatment requires 6 mL for each side of the face using a 26-gauge needle.

TABLE 1

Growth in the Use of Soft Tissue Fillers: 2005 to 2006¹

	Procedures, n		
Soft Tissue Filler	2005	2006	Change, %
Hyaluronic acid	489,554	778,285	59
Collagen	220,632	267,339	21
Calcium hydroxylapatite	66,182	78,849	19
Poly-L-lactic acid	46,732	54,912	18
Fat	48,960	52,904	8
Total	872,060	1,232,289	41

The use of poly-L-lactic acid can be accompanied by the delayed occurrence of subcutaneous papules at the injection site. These are typically palpable, asymptomatic, and nonvisible. Other treatment-related adverse events include bruising, edema, hematoma, inflammation, and erythema.³

Nasolabial and Mesolabial Folds

For grade 3 or 4 (mild to moderate) nasolabial folds, calcium hydroxylapatite is used. As opposed to poly-L-lactic acid, in which the full response requires time for collagen deposition, the clinical response to calcium hydroxylapatite is related to injection volume.

TABLE 2

The 9 Most Commonly Offered Noninvasive Procedures by the American Academy of Cosmetic Surgery Member Practices in 2006*2

Procedure	Member Practices Offering Procedure, %
Botulinum toxin type A	94
Hyaluronic acid	92
Chemical peels	80
Laser resurfacing	69
Microdermabrasion	65
Fat injections	62
Collagen injections	59
Laser hair removal	59
Calcium hydroxylapatite/poly-L-lactic acid [†]	50
*Based on 206 completed surveys	

^{*}Based on 206 completed surveys.

[†]The American Academy of Cosmetic Surgery confirms that calcium hydroxylapatite/poly-L-lactic acid data are combined.

TABLE 3

The 5 Surgical Procedures Offered More Commonly Than Calcium Hydroxylapatite/Poly-L-lactic Acid by American Academy of Cosmetic Surgery Member Practices in 2006²

Procedure	Member Practices Offering Procedure, %
Liposuction	76.2
Blepharoplasty	68.4
Face-lift	59.7
Forehead-lift	58.3
Lip implant	51.9

Over time, the manufacturer claims that calcium hydroxylapatite also stimulates new collagen growth. Clinical results last for 9 to 12 months.⁴

The treatment of more severe nasolabial and mesolabial folds may require injections of hyaluronic acid, with additional filling accomplished using underlying poly-L-lactic acid injections.

Tear Troughs

Individuals in their early 30s begin to experience some descent of the malar fat pad. This can result in the formation of dark circles beneath the eyes and deepening of the nasolabial and nasojugal creases.⁵

The hyaluronic acid formulation in Juvéderm differs from that of Restylane in that there is less resistance to injecting the product through the syringe. However, based on early experience with Juvéderm, it is important to secure the needle firmly to the syringe in order to avoid separation during injection.

The most common side effects from hyaluronic acid include temporary injection site reactions such as redness, pain and tenderness, firmness, swelling, lumps and bumps, and bruising.

LOWER FACE

Most patients want to add volume to their lips but wish to avoid looking overdone. In addition, a common complaint among women older than 50 years is lipstick "bleeding" around the mouth along the perioral rhytides.⁶

Lips

My preference is to use Perlane, a formulation of hyaluronic acid. The objective is to provide a more youthful and relaxed appearance with an uplifting hint of a smile. This is accomplished using injections at the corners of the mouth (oral commissure) at the level of the modiolus. Additionally, injections along the philtral columns enhance and redefine the Cupid's bow area. Successful treatment in this anatomical area requires attention to facial characteristics and ethnicity.^{7,8}

Perlane is supplied in a 1-mL disposable glass syringe. The most commonly observed side effects are swelling, redness, pain, bruising, and tenderness at the injection site, which typically resolve in fewer than 7 days.⁹

Upper- and Lower-Lip Rhytides

A human-based collagen product, CosmoDerm, can be used to correct upper- and lower-lip rhytides. Unlike hyaluronic acid, calcium hydroxylapatite, and poly-L-lactic acid, human-based collagen implants are sterile injectable liquids made of highly purified human collagen. These products (ie, CosmoDerm, CosmoPlast) also contain lidocaine to provide local anesthesia at the site of injection. ¹⁰

Human-based collagen implants can be injected without a skin test prior to treatment. However, they should not be used in patients with severe or multiple allergies that have led to anaphylactic shock or an acute reaction that requires immediate emergency medical assistance. Human-based collagen implants should also not be used in patients who are allergic to lidocaine or in patients with an inflamed or infected skin condition.¹⁰

SUMMARY

As the availability of new products for nonsurgical facial rejuvenation continues to grow, we can expect

CHOOSING APPROPRIATE FILLERS

that more clinicians will utilize these new products in order to meet their patients' expectations for a youthful appearance. The recommendations in this article should not be interpreted as rigid rules. Rather, they should stimulate personal consideration of the options available to enhance the practice of dermatologists and plastic surgeons. Ultimately, becoming skilled in the use of several categories of fillers will provide the best outcomes.

REFERENCES

- 1. American Society of Plastic Surgeons. 2000/2005/2006 National Plastic Surgery Statistics: Cosmetic and Reconstructive Procedure Trends. http://www.plasticsurgery.org/media/statistics/loader.cfm?url=/commonspot/security/getfile.cfm&PageID=23628. Accessed December 19, 2007.
- American Academy of Cosmetic Surgery 2006 Procedural Census. Prepared by RH Research. 2006.
- 3. Sculptra [package insert]. Berwyn, PA: Dermik Laboratories; 2004.

- 4. Radiesse [package insert]. Franksville, WI: BioForm Medical; 2006.
- Robertson KM, Ramirez O. Facelift, mid-face. October 3, 2006. http://www.emedicine.com/plastic/TOPIC47.htm. Accessed November 19, 2007.
- Beer KR. Rejuvenation of the lip with injectables. July 12, 2007. http://www.medscape.com/viewarticle/559079_1. Accessed November 19, 2007.
- Werschler WP. Combining advanced injection techniques: poly-L-lactic acid as the foundation for nonsurgical total facial rejuvenation and restoration. *Cosmet Dermatol.* 2007;20(suppl 1):9-13.
- 8. Dev VR, Wang P. Lip reduction. June 14, 2006. http://www.emedicine.com/plastic/topic66.htm. Accessed November 28, 2007.
- 9. Perlane [package insert]. Scottsdale, AZ: Medicis Aesthetics; 2007.
- 10. US Food and Drug Administration Web site. New device approval: CosmoDerm™ 1 Human-Based Collagen, CosmoDerm™ 2 Human-Based Collagen and CosmoPlast™ Human-Based Collagen P800022/S050. http://www.fda.gov/cdrh/mda/docs/p800022s050.html. Accessed November 19, 2007.

Dermal Fillers for Facial Rejuvenation and Restoration: Integrating New Therapies Into Clinical Practice

This activity has been certified for physicians. It was planned and produced in accordance with the ACCME essentials and standards for enduring materials (release date: February 2008; expiration date: February 2009). To obtain CME credit, please complete this form, remove from the booklet, and return to Postgraduate Institute for Medicine at 367 Inverness Parkway, Suite 215, Englewood, CO 80112, or fax to 303-790-4876.

For each question listed below, select the one best answer.

- 1. What are the applications for facial-shaping agents?
 - a. Enhance natural features
 - b. Restore aged facial features
 - c. Rejuvenate fading youth
 - d. All of the above
- 2. Select the volumetric lifting procedure that serves as an alternative and a complement to traditional surgical tension lifting.
 - a. Regional aesthetic volume enhancement
 - b. Nonsurgical total facial rejuvenation and restoration
 - c. Both a and b
 - d. None of the above
- 3. Which answer best describes regional aesthetic volume enhancement?
 - a. A method of assessing, planning, and performing volume enhancement to replace (augment) biometric volume loss/alteration
 - b. A surgical technique to provide long-term lifting of aging facial features
 - c. A product-dependent procedure that provides volumetric lifting of facial features
 - d. All of the above
- 4. Select the term that best describes hyaluronic acid.
 - a. Stimulatory filler
 - b. Chemical denervation
 - c. Replacement filler
 - d. Both a and c
- 5. Select the characteristic finding(s) associated with intrinsic aging.
 - a. Slowed skin and collagen production
 - b. Changes in elastin resulting in a loss of tone
 - c. Sun exposure
 - d. Both a and b

- 6. Identify the products composed of humanderived collagen.
 - a. Captique, Hylaform, Juvéderm, and Restylane
 - b. CosmoDerm and CosmoPlast
 - c. Radiesse and Sculptra
 - d. Both a and b
- 7. Silicone and polymethylmethacrylate are synthetic products that are not readily metabolized and removed by the body.
 - a. True
 - b. False
- 8. According to the American Society of Plastic Surgeons survey, what was the growth in the use of all fillers from 2005 to 2006?
 - a. 14%
 - b. 24%
 - c. 41%
 - d. 91%
- 9. According to the American Academy of Cosmetic Surgery, how many of its members offer semipermanent fillers to their patients?
 - a Less than 10%
 - b. Approximately one quarter
 - c. Approximately one half
 - d. More than 9 of 10
- 10. Select the correct statement regarding the hyaluronic acid products Juvéderm and Restylane.
 - a. Greater resistance injecting Juvéderm through the syringe
 - b. Less resistance injecting Juvéderm through the syringe
 - c. Severe nasolabial folds might require additional filling with poly-L-lactic acid
 - d. Both a and c

APPLICATION FOR CATEGORY I CREDITS

Dermal Fillers for Facial Rejuvenation and Restoration: Integrating New Therapies Into Clinical Practice

Project ID: 4991ES22 Released: February 2008

To assist us in evaluating the effectiveness of this activity and to make recommendations for future educational offerings, please take a few minutes to complete this evaluation form. You must complete this evaluation form to receive acknowledgment for completing this activity.

Please answer the followir 1=Strongly Disagree	ng questions by circ 2=Disagree	cling the appropriat 3=Neutral	e rating: 4=Agree	5=Strongly Agree
Extent to Which Progra	m Activities Met	the Identified Ob	jectives	G, G
After completing this activity	y, I am now better al	ole to:		
• Define the techniques of				1 2 3 4 5
• Identify a personal plan				
 Describe the use of colla and restoration 	igen stimulators for	nonsurgical treatn	nent of facial rejuv	
 Identify treatment optio 	ne bacad on locatio	on and stage of line	atrophy	1 2 3 4 5
 Describe safety consider 				1 2 3 4 5
Overall Effectiveness of		ii teeiiiiques ioi ue	mai micis	1 2 3 , 3
The content presented:	the Activity			
• Was timely and will infl	uence how I praction	ce		1 2 3 4 5
• Enhanced my current ki	nowledge base			1 2 3 4 5
 Addressed my most pres 	ssing questions			1 2 3 4 5
 Provided new ideas or in 	nformation I expect	t to use		1 2 3 4 5
 Addressed competencies 	s identified by my s	specialty		1 2 3 4 5
 Avoided commercial bia 	s or influence			1 2 3 4 5
Impact of the Activity				
Name one thing you inter	d to change in you	ir practice as a resu	lt of completing th	nis activity:
Additional comments abo	ut this activity:			
ollow-up				
s part of our continuous quucational interventions or	n professional pract	tice. Please indicate	if you would be w	w-up surveys to assess the impact villing to participate in such a sur sted in participating in a follow-up su
2 3	4 5	_ 6 7	8 9	10
equest for Credit				
ame:			Degree:	
rganization:				
ldress:				
ty:				Zip Code:
elephone:				E-mail:
1				
_				
or Physicians Only certify my actual time sper I I participated in the entire ac				t of the activity and claim cred



A supplement to Cosmetic Dermatology®

Dermal Fillers for Facial Rejuvenation and Restoration: Integrating New Therapies Into Clinical Practice

This activity is jointly sponsored by Postgraduate Institute for Medicine and The MedCom Resource, Inc.

Supported by an educational grant from Dermik Laboratories, a business of sanofi-aventis U.S. LLC.