

# REVANESSE®



2021 NEXT GENERATION HA | Physician Presentation

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- Versa™

- **REAL PEOPLE REAL RESULTS**



PROLLENIUM®

# WHO IS PROLLENIUM®?

- **Founded in 2002**, Prollenium® is an aesthetic medical device company
- **Only manufacturer** of dermal fillers in North America
- Heavily invested in research and development
- Reshaping the aesthetic experience

For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

The Prollenium logo is displayed in a stylized, glowing font. The letter 'O' is replaced by a circular graphic element consisting of concentric arcs, giving it a modern, high-tech appearance.



# MANUFACTURED IN CANADA

## TWO STATE-OF-THE-ART MANUFACTURING FACILITIES

- **Highest quality** manufacturing and QA standards
- **Highly regulated** adhering to Good Manufacturing Practices (GMPs)
- **Regulatory compliance** to the highest standards (ISO 13485:2016)
- **FDA approved** facility

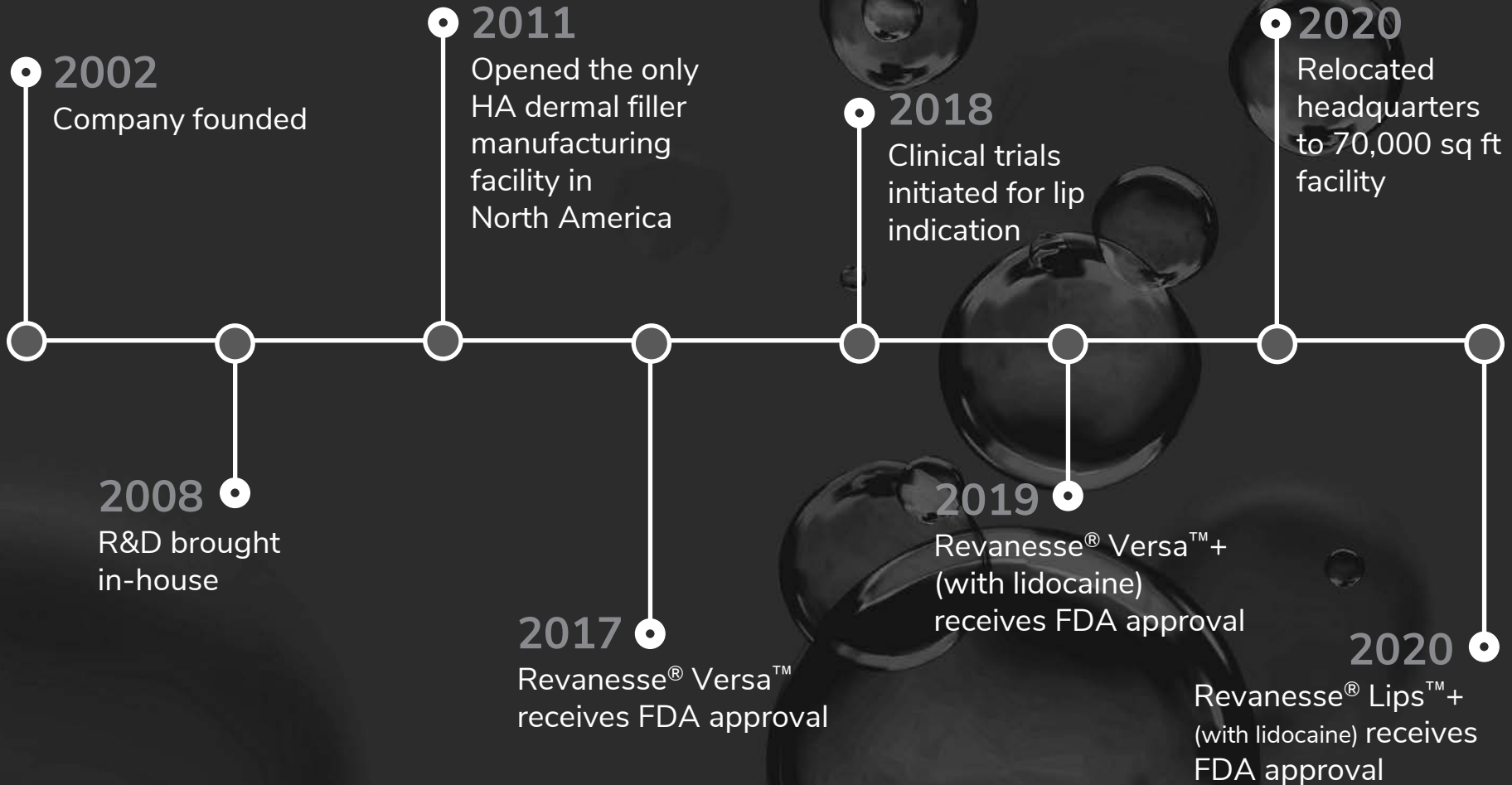


# CONSISTENCY

**OUR MANUFACTURING  
AND QUALITY  
ASSURANCE PROCESSES  
CREATE PREDICTABLE  
RESULTS FOR INJECTOR  
CONFIDENCE AND  
PATIENT SAFETY**



# COMPANY TIMELINE



For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](https://www.RevanesseUSA.com)





# Story of Revanesse®



# WHAT IS REVANESSE® ?

- Revanesse® is a line of injectable Hyaluronic Acid (HA) dermal fillers that use state-of-the-art production methods, including proprietary cross-linking and shaping processes that yield uniform, spherical particles.
- Revanesse® Versa™ is indicated for injection into the mid to deep dermis for correction of moderate to severe facial wrinkles and fold, including nasolabial folds in adults 22 years of age or older.

For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

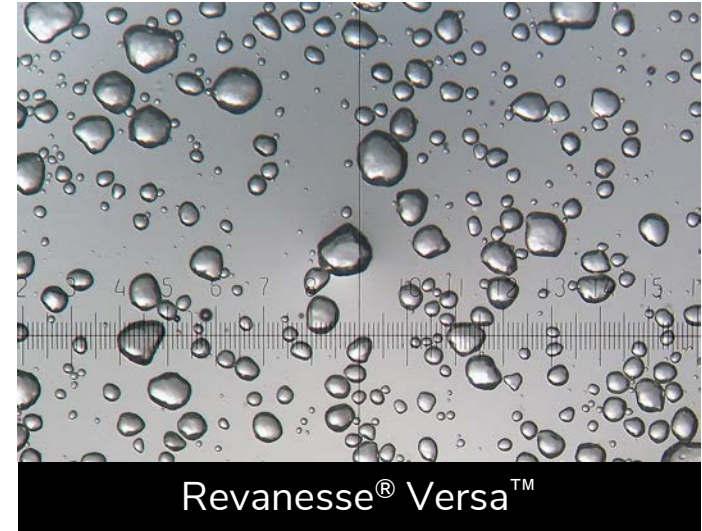


# UNIQUE FORMULATION

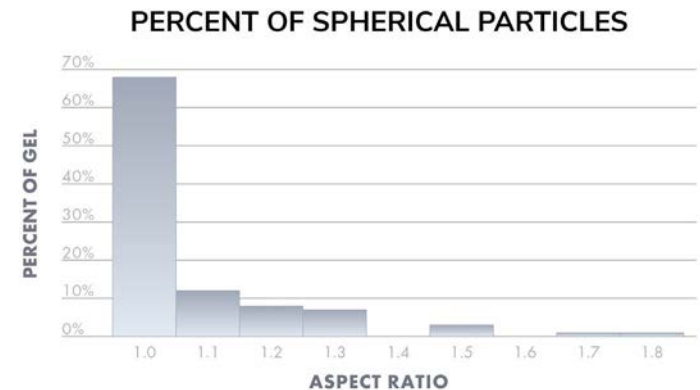
- **Proprietary** sieving technology produces round, smooth and spherical HA particles<sup>1</sup>
- **Optimally** cross-linked, long-chain, high molecular weight (25mg/mL) HA using less BDDE
- **7-Day Dialysis** to neutralize pH and remove excess BDDE
- **Rigorous** quality control throughout manufacturing process

1. BDDE Cross-linked Hyaluronan Dermal Fillers Comparison of Commercial Products Update Report RD045, 2014. Prolenium Medical Technologies Inc.

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20X Magnification

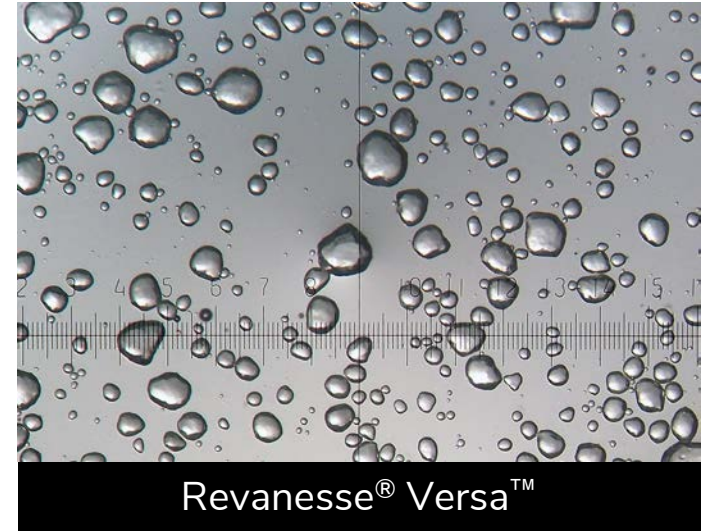


Test batch of Revanesse® Versa™. Graph demonstrates 68% perfectly spherical particles, where the aspect ratio of a perfect sphere is 1.0.



# DESIGNED FOR SAFETY AND LONGEVITY

- **Reduced inflammatory reaction:** Inflammatory reactions are more pronounced for particles of irregular shape<sup>1</sup>
  - Revanesse® uses only high molecular weight HA to minimize the inflammatory response for tissue integration<sup>2</sup>
  - Round and smooth HA particles that use high molecular weight HA are less likely to be recognized as foreign bodies<sup>1,3</sup>
- **Durability and long-lasting:** Degree of cross-linking is optimized using Thixofix® technology.
  - Unique sieving process helps produce uniform and spherical particles
- **Optimized for:**
  - Elasticity (G') for lift capacity<sup>1</sup>
  - Yield stress for shape retention<sup>1</sup>
  - Cohesivity for tissue integration<sup>1</sup>



20X Magnification

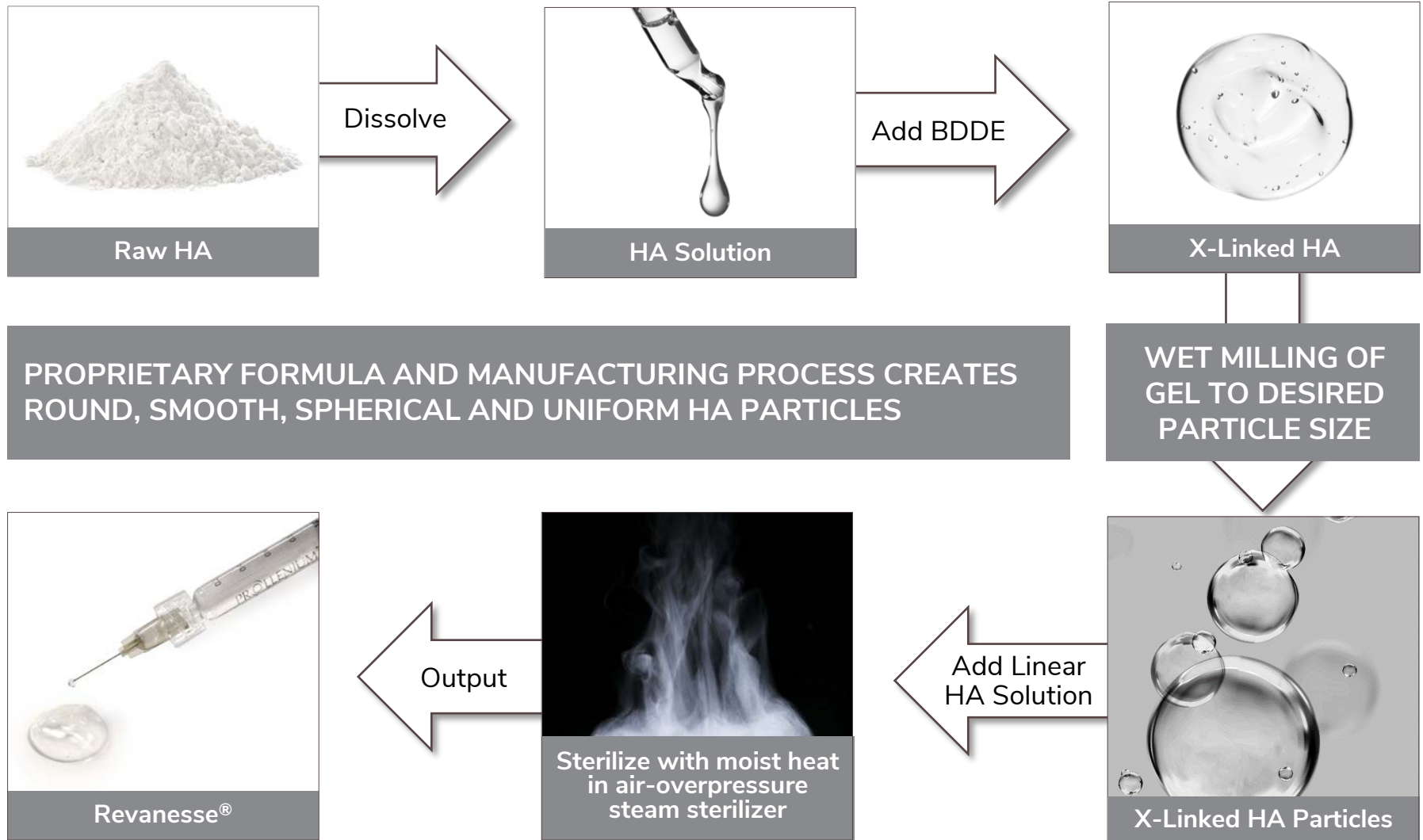
1. Biocompatibility of Microparticles into Soft Tissue Fillers. *Semin Cutan Med Surg.* 2004; 23(4): 214-217.
2. Testing from Prollenium Medical Technologies Inc. Data on file.
3. High and Low Molecular Weight Hyaluronic Acid Differentially Influence Macrophage Activation. *ACS Biomater Sci Eng.* 2015; 1(7): 481-493.

For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

A black and white photograph showing a medical syringe with a needle. The needle is positioned just above a large, clear droplet of liquid that has already been deposited on a light-colored surface. A smaller droplet is still attached to the tip of the needle. The syringe has markings on its barrel, including a 'CE' symbol and some numbers. The background is out of focus.

**Manufacturing Process**

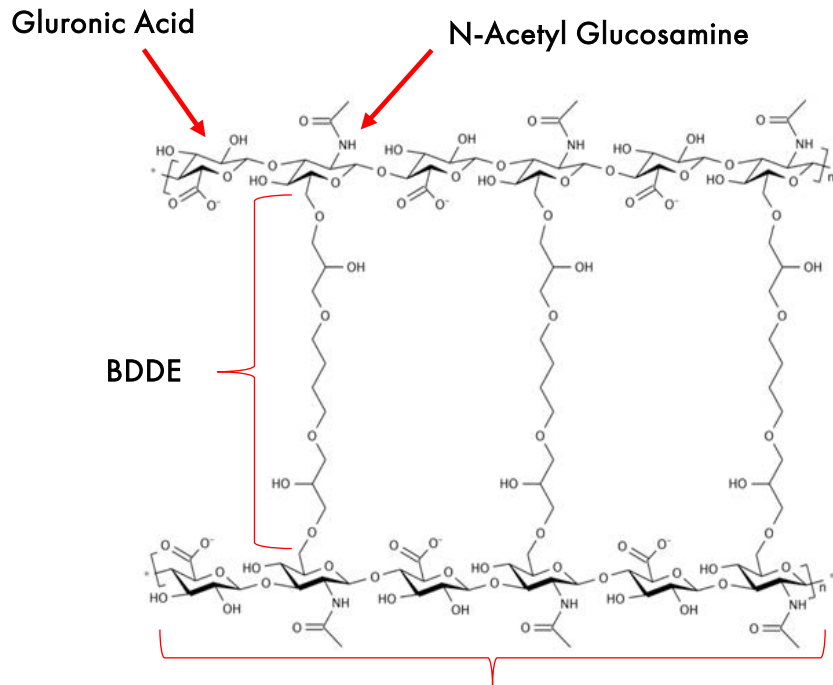
# PRODUCT SYNTHESIS FLOW CHART



For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

# MODIFICATION

THE EFFECT OF MODIFICATION IS TO INCREASE DURABILITY (STABILITY) & LONGEVITY



Hyaluronic Acid (HA)

- Natural Polysaccharide
- Ubiquitous
- 15 g / person

Modification helps promote cross-links between different HA polymer chains.

Excessive cross-linking can create hard implants and increase the safety risk or incidence of adverse reactions (AEs).

Optimized modification is important for balancing safety and durability.

Testing from Prolenium Medical Technologies Inc. Data of file.

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# MOLECULAR WEIGHT

REVANESSE® EXCLUSIVELY USES HIGH MOLECULAR WEIGHT HYALURONIC ACID WHICH REQUIRES LESS BDDE TO MAKE EFFECTIVE LINKS

Less BDDE is required to link higher molecular weight HA

- Reduces incidence of inflammatory response

Shorter HA chains have lower molecular weight and require more BDDE to achieve effective links

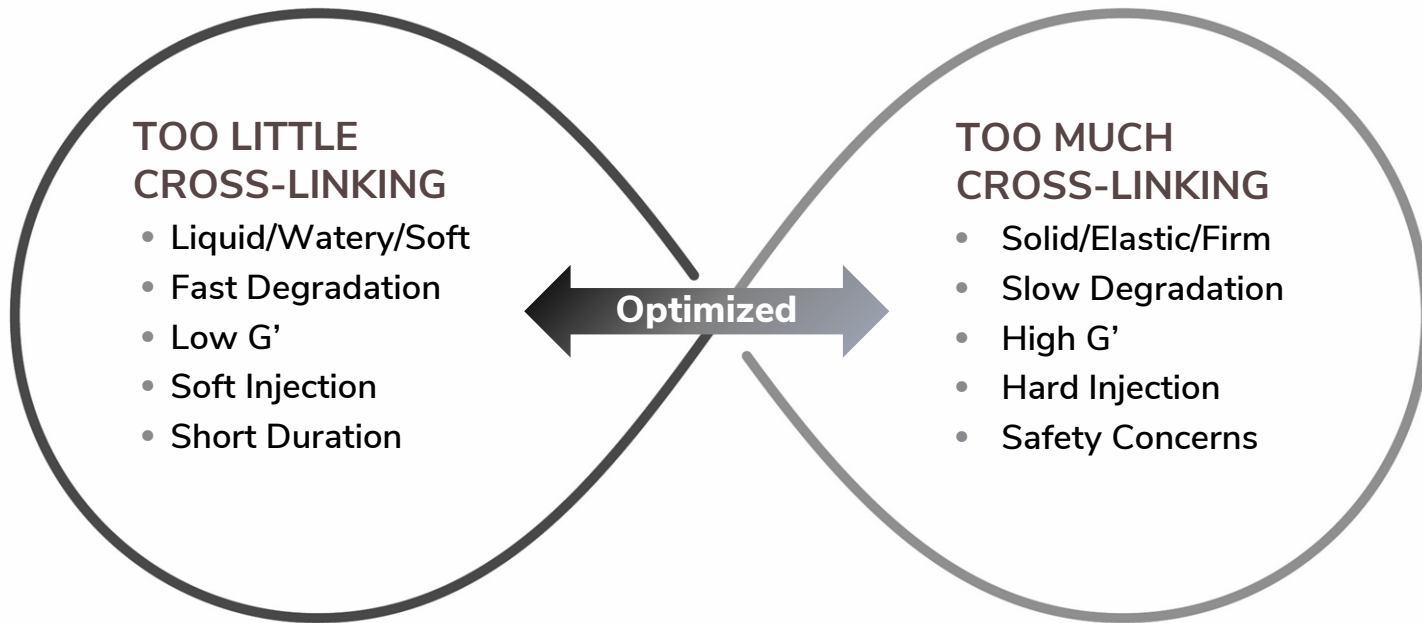
- Low and medium molecular weight HA may potentially increase inflammatory responses
- Using more BDDE theoretically makes modified HA less natural and more likely to produce adverse reactions

High and Low Molecular Weight Hyaluronic Acid Differentially Influence Macrophage Activation. *ACS Biomater Sci Eng.* 2015; 1(7): 481-493.

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# OPTIMIZATION

REVANESSE USES THIXOFIX® TECHNOLOGY TO OPTIMALLY CROSS-LINK HIGH MOLECULAR WEIGHT HA

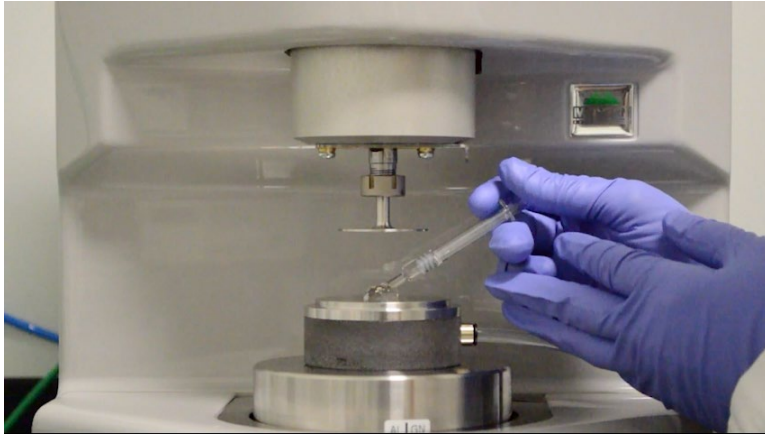


Using proprietary Thixofix® cross-linking technology, Revanesse® Versa™ has a balance between safety and duration

For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

# ELASTICITY ( $G'$ )

$G'$  IS THE MEASURE OF ELASTICITY AND CALLED THE ELASTIC MODULUS



Rheometer



Jello-Like (Elastic)

Measured via a rheometer

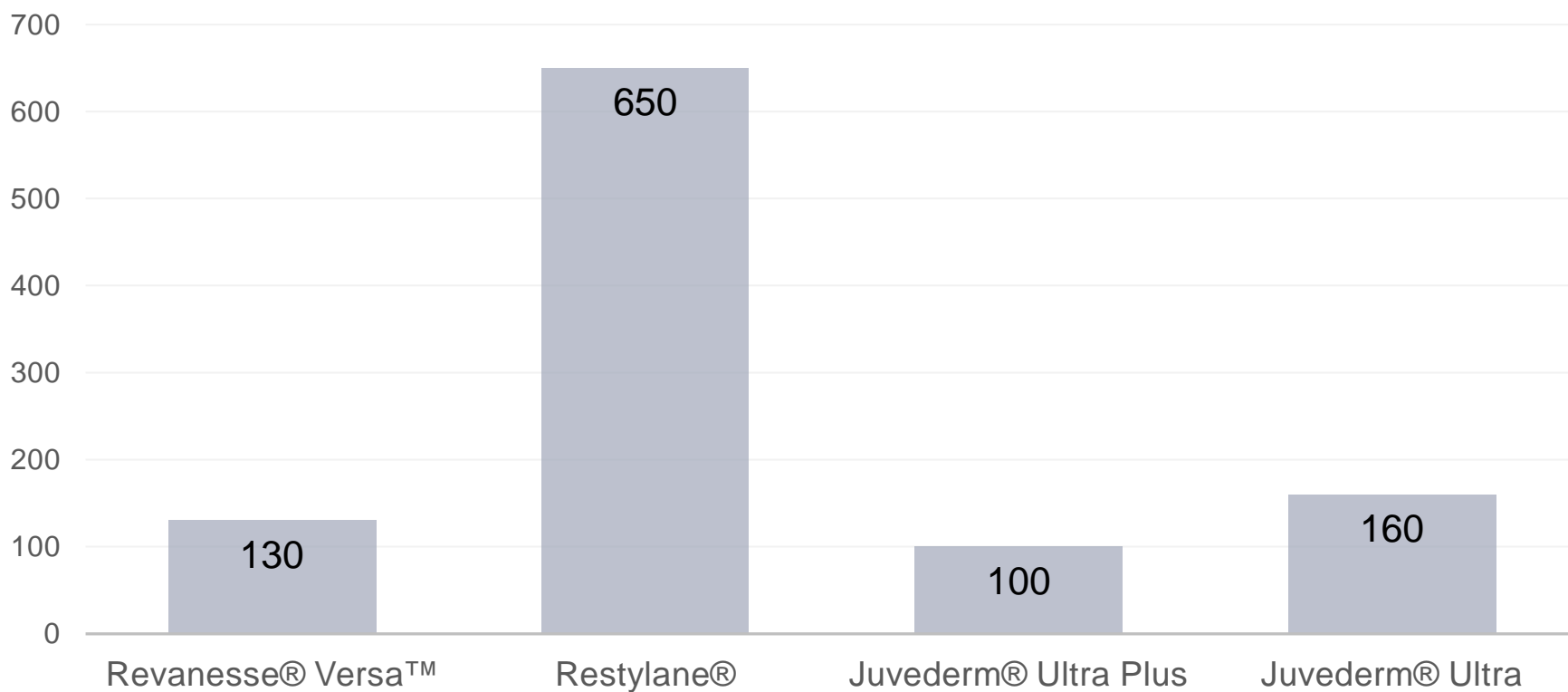
$G'$  establishes the elastic behavior (stiffness or lift capacity) of an HA gel or its ability to recover its shape after shear deformation (injection process)

For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)



# G' ELASTIC MODULUS (1HZ)

G' IS THE MEASURE OF ELASTICITY AND CALLED THE ELASTIC MODULUS



BDDE Cross-linked Hyaluronan Dermal Fillers Comparison of Commercial Products Update Report RD045  
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# VISCOSITY

VISCOSITY IS A MEASURE OF RESISTANCE TO FLOW (THICKNESS) WHEN SHEAR DEFORMATION IS APPLIED DURING THE INJECTION PROCESS

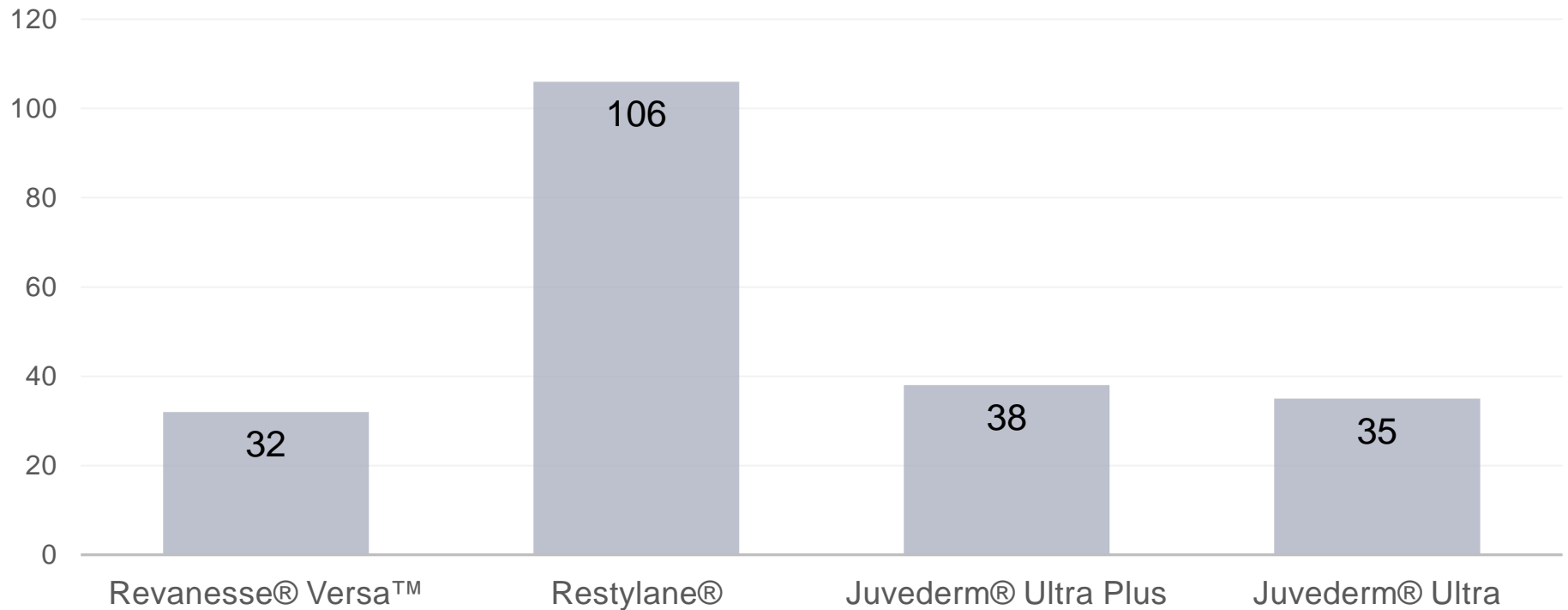


Honey-Like (Flow)

Viscosity measures how liquid the product is for creating volume

# G'' VISCOUS MODULUS (0.7HZ)

VISCOSITY IS A MEASURE OF RESISTANCE TO FLOW (THICKNESS) WHEN SHEAR DEFORMATION IS APPLIED DURING THE INJECTION PROCESS

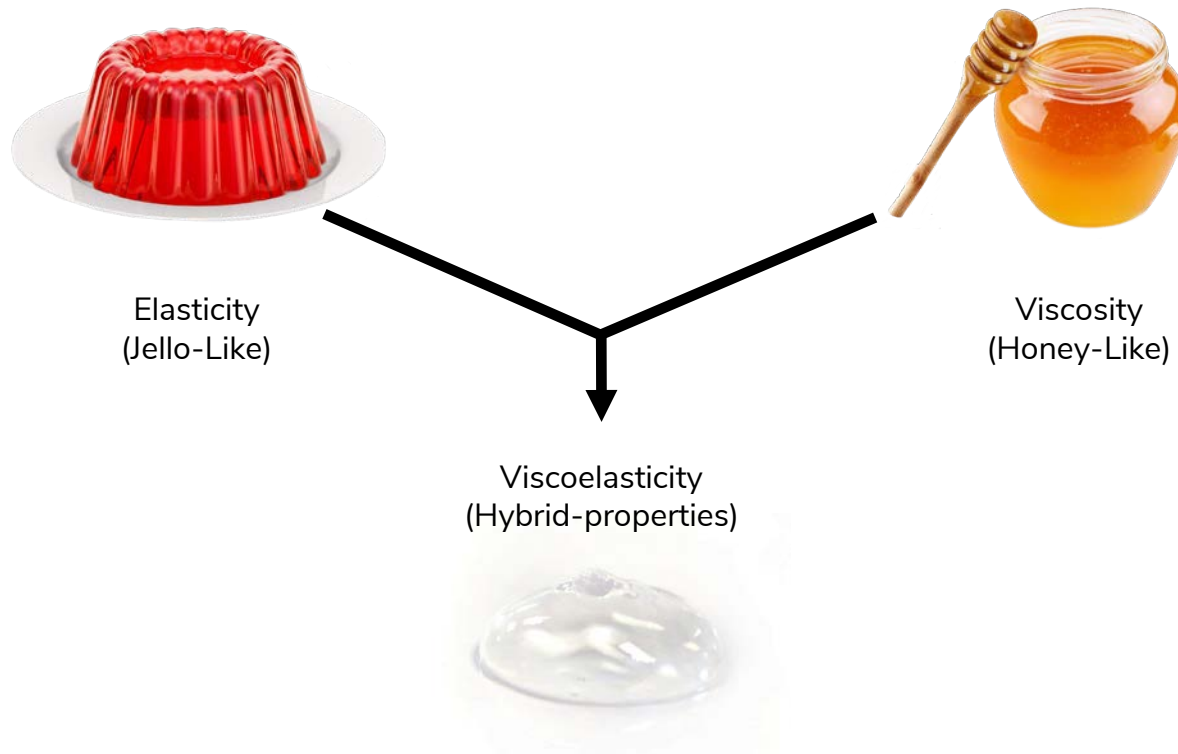


M.H. Gold, Stafford Baumann, C.P. Clark III, J. Schlessinger

For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

# VISCOELASTICITY

REVANESSE® PRODUCTS OFFER THE NECESSARY VISCOELASTICITY



For smooth extrusion  
during injections

Ideal mechanical  
properties after injections

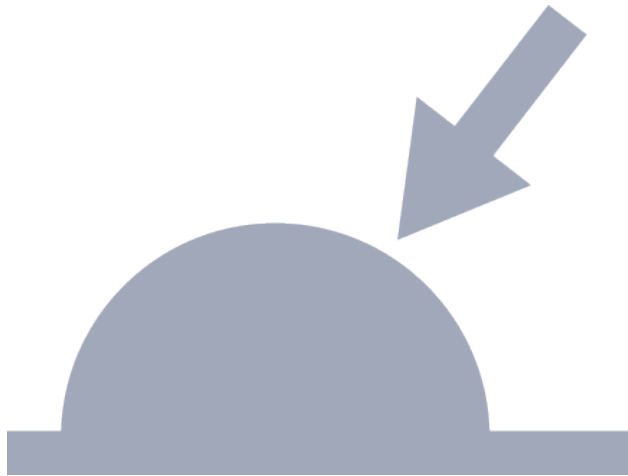
For treating different  
injectable depths

For adapting to dynamic  
facial forces

Testing from Prollenium Medical Technologies Inc. Data of file.  
For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

# YIELD STRESS

YIELD STRESS IS A MEASURE OF THE AMOUNT OF FORCE REQUIRED TO DEFORM THE HA GEL



Shape Retention

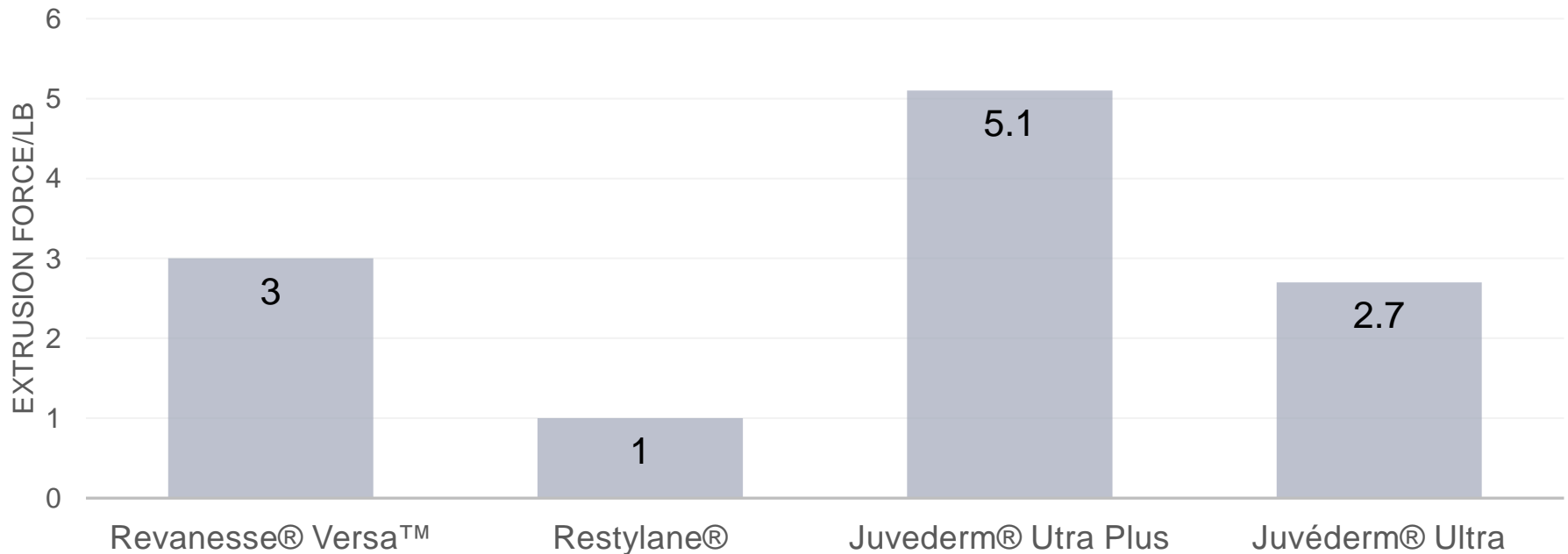
Provides insight into how well an HA gel deposit retains its initial shape after implantation

Testing from Prollenium Medical Technologies Inc. Data of file.

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# G'' VISCOUS MODULUS (0.7HZ)

VISCOSITY IS A MEASURE OF RESISTANCE TO FLOW (THICKNESS) WHEN SHEAR DEFORMATION IS APPLIED DURING THE INJECTION PROCESS



BDDE Cross-linked Hyaluronan Dermal Fillers Comparison of Commercial Products Update Report RD045  
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# PRODUCT OVERVIEW

Test	Versa™	Benefit
Needle (G)	27	Size of the needle bore for determining how much product is extruded
Degree of Modification – % modified w/ BDDE	6.9%	Higher amount increases stability for durability and longevity
Mean Extrusion Force (LBS)	2.4	Provides injection control for accuracy and precision
G' (Pa @ 1.0Hz)	110	Creates stiffness (lift capacity); measures how <u>solid</u> it is
Degradation Rate (Pa/H)	2.2	Duration and safety; higher number results in faster absorption
Viscosity (Pa.S) – <u>not</u> G"	2500	Assists in creating volume; measures how <u>liquid</u> it is
Swell Ratio (SwF) – To Reach Equilibrium	3.7X	Low water absorption after implantation
Cohesivity (mg/drop) – Drop Weight Method	26	How well the gel resists dynamic facial forces after implantation; higher value provides more resistance
Yield Stress (Pa) - Shape Retention	23.6	How the product retains its shape; how much force is required to deform the gel

Testing from Prollenium Medical Technologies Inc. Data of file.

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# Versa™ vs Nasolabial Fold Comparator

# STUDY DESIGN

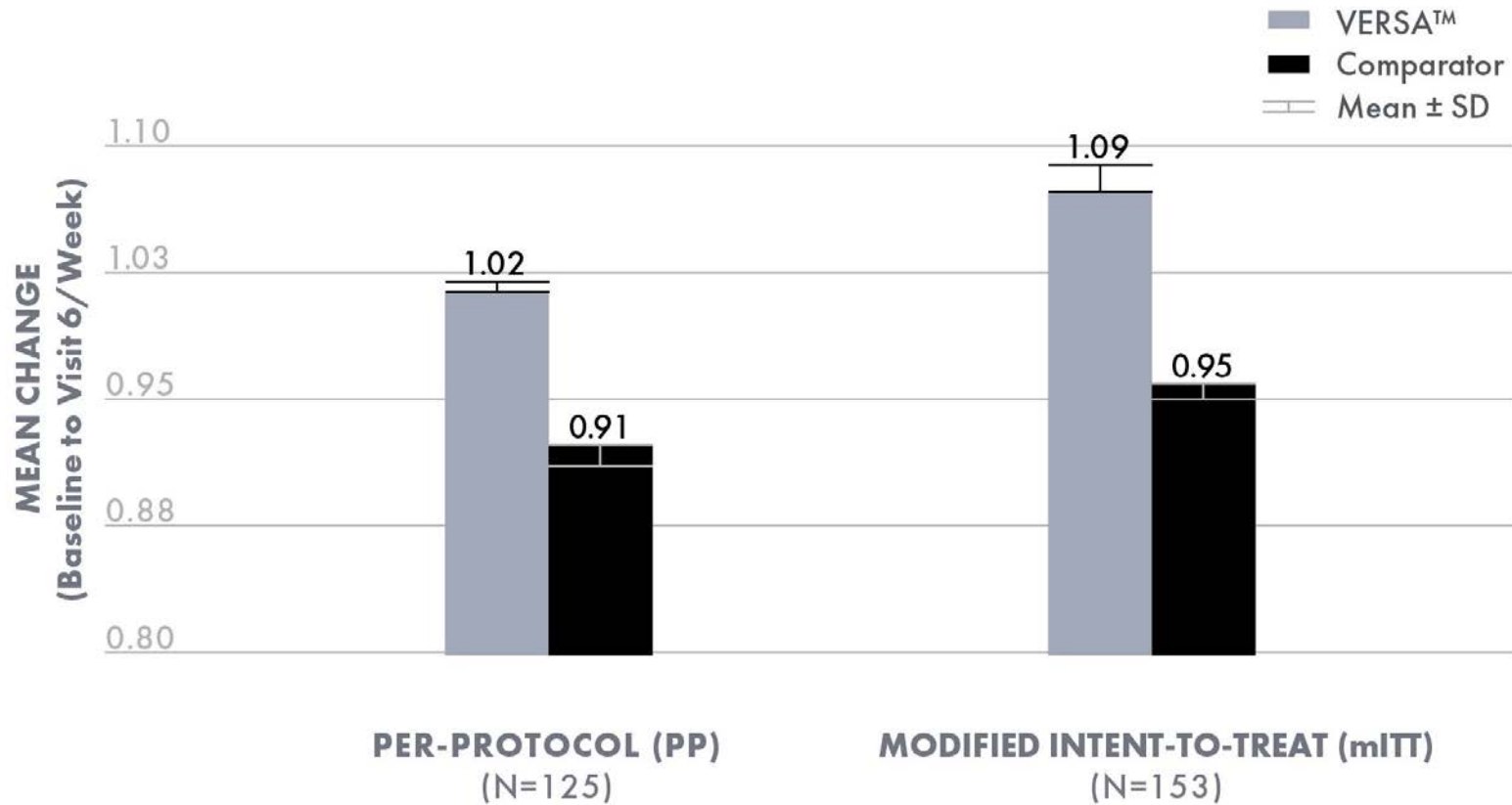
## A MULTICENTER, DOUBLE-BLIND, RANDOMIZED, SPLIT-FACE STUDY TO EVALUATE THE SAFETY AND EFFICACY OF REVANESSE® VERSA™ VS COMPARATOR FOR THE CORRECTION OF NASOLABIAL FOLDS

- Qualified subjects had Nasolabial Folds (NLFs) with a Wrinkle Severity Rating Scale (WSRS) score of 3 or 4 (moderate or severe)
- Side by side comparison: NLFs treated with Revanesse® Versa™ on one side of the face and Comparator on the other side of the face
- Side of the face for each product was randomly assigned
- Evaluating investigator and subject were blinded and injections were performed by unblinded physician
- Maximum of 2mL per fold
- All initial treatments were administered at baseline and in addition to WSRS, evaluations included the Global Aesthetic Improvement scale (GAI) of the investigator and the patients, as well as adverse events recorded in a diary of each subject
- Based on use of photographs, the WSRS is designed to quantify facial folds by visual assessment of the length and apparent depth of the fold without referring to baseline
- In contrast, the GAI scale is used to grade overall improvement in each fold by comparing its appearance at follow up against a high magnification photograph taken before treatment
- For subjects not requiring retreatment, the study period ended at Visit 6 / Week 24

Revanesse® Versa™ Clinical Study Report SYM 2014-02: A Multicenter, Double-Blind, Randomized, Split-Face Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ versus Restylane® for the Correction of Nasolabial Folds. Revanesse® Versa™ is a registered trademark of Prollenium Medical Technologies Inc. Restylane® is a registered trademark of Nestlé Skin Health S.A. For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

# 1° EFFICACY

## MEAN WRINKLE SEVERITY RATING SCALE (WSRS)

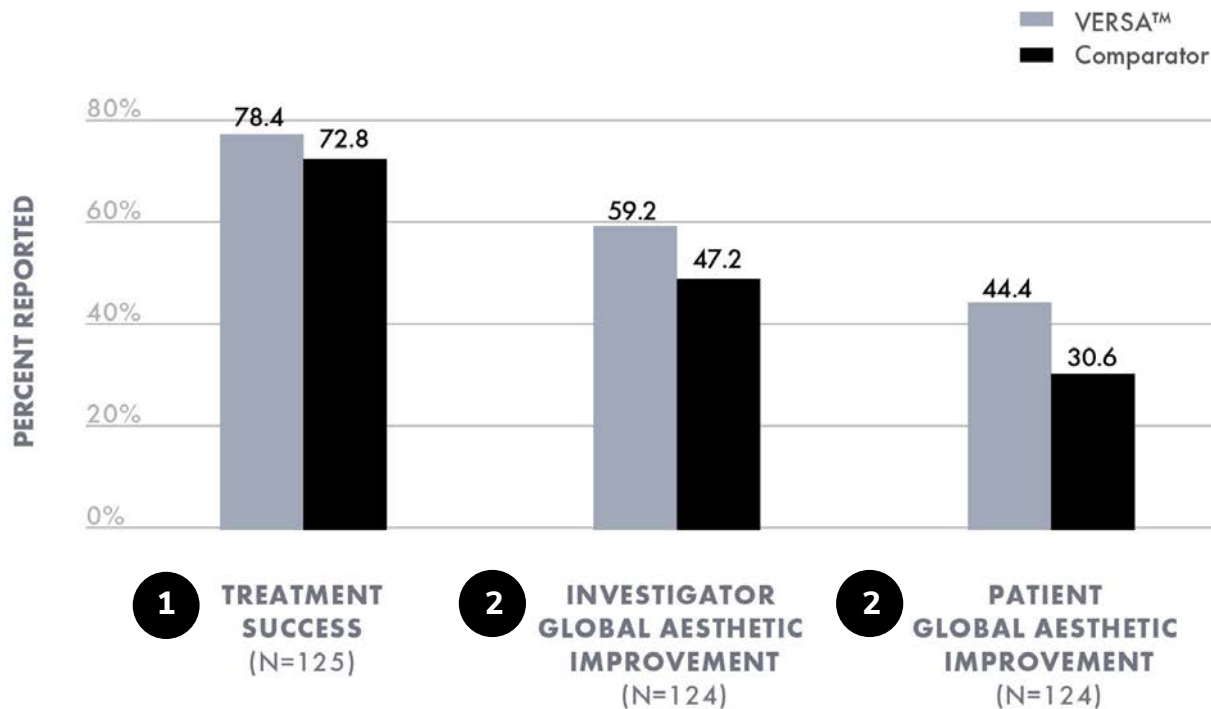


Revanesse® Versa™ Clinical Study Report SYM 2014-02: A Multicenter, Double-Blind, Randomized, Split-Face Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ versus Restylane® for the Correction of Nasolabial Folds. Revanesse® Versa™ is a registered trademark of Prolenium Medical Technologies Inc. Restylane® is a registered trademark of Nestlé Skin Health S.A.

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# 2° EFFICACY

## PERCENT AT VISIT 6 / WEEK 24 (PER-PROTOCOL)



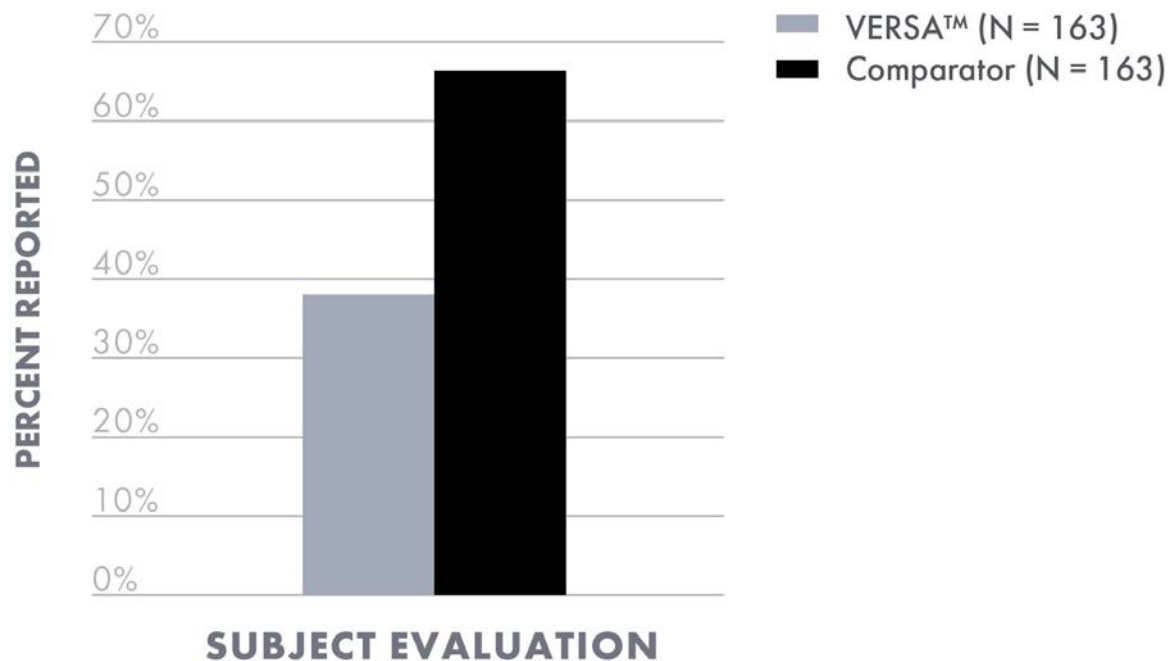
- 1 At least a 1-grade improvement in WSRS score from baseline
- 2 Categories of “much improved” or “very much improved”

Revanesse® Versa™ Clinical Study Report SYM 2014-02: A Multicenter, Double-Blind, Randomized, Split-Face Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ versus Restylane® for the Correction of Nasolabial Folds. Revanesse® Versa™ is a registered trademark of Prollenium Medical Technologies Inc. Restylane® is a registered trademark of Nestlé Skin Health S.A.

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# TREATMENT EMERGENT ADVERSE EVENTS

## PAIN AT INJECTION SITE



66.3% of the ITT subjects treated with Comparator reported pain at the injection site vs. 38.0% of the subjects treated with Versa™

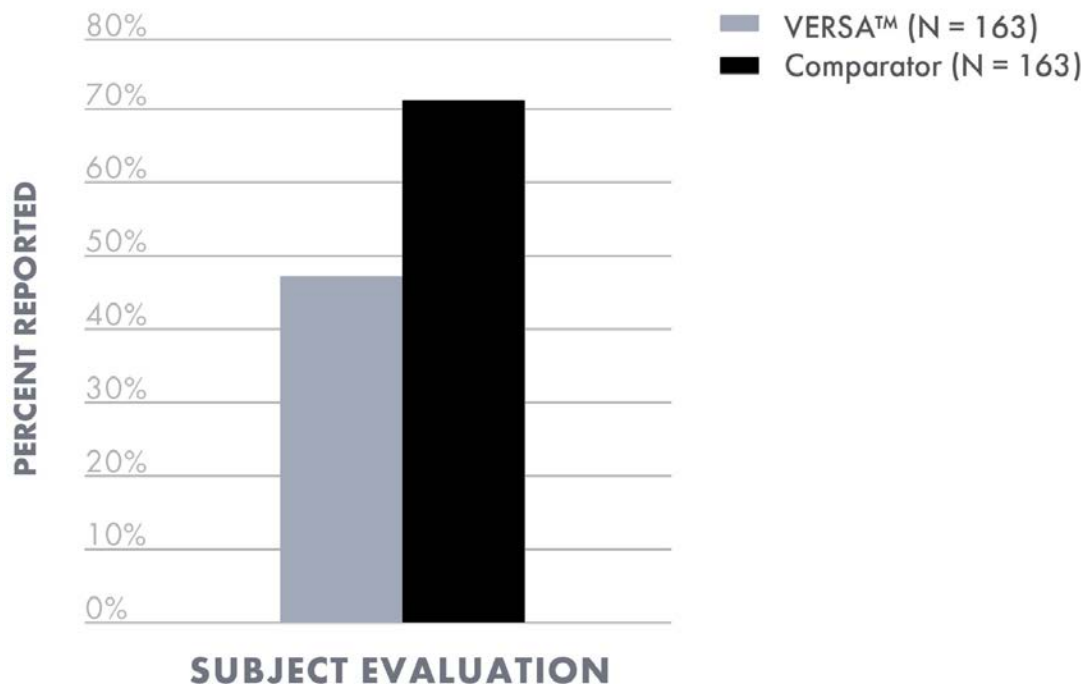
Revanesse® Versa™ Clinical Study Report SYM 2014-02: A Multicenter, Double-Blind, Randomized, Split-Face Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ versus Restylane® for the Correction of Nasolabial Folds. Revanesse® Versa™ is a registered trademark of Prollenium Medical Technologies Inc. Restylane® is a registered trademark of Nestlé Skin Health S.A.

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ITT: Intent-to-Treat subjects

# TREATMENT EMERGENT ADVERSE EVENTS

## SWELLING AT INJECTION SITE



71.2% of the ITT subjects treated with Comparator reported swelling at the injection site vs. 47.2% of the subjects treated with Versa™

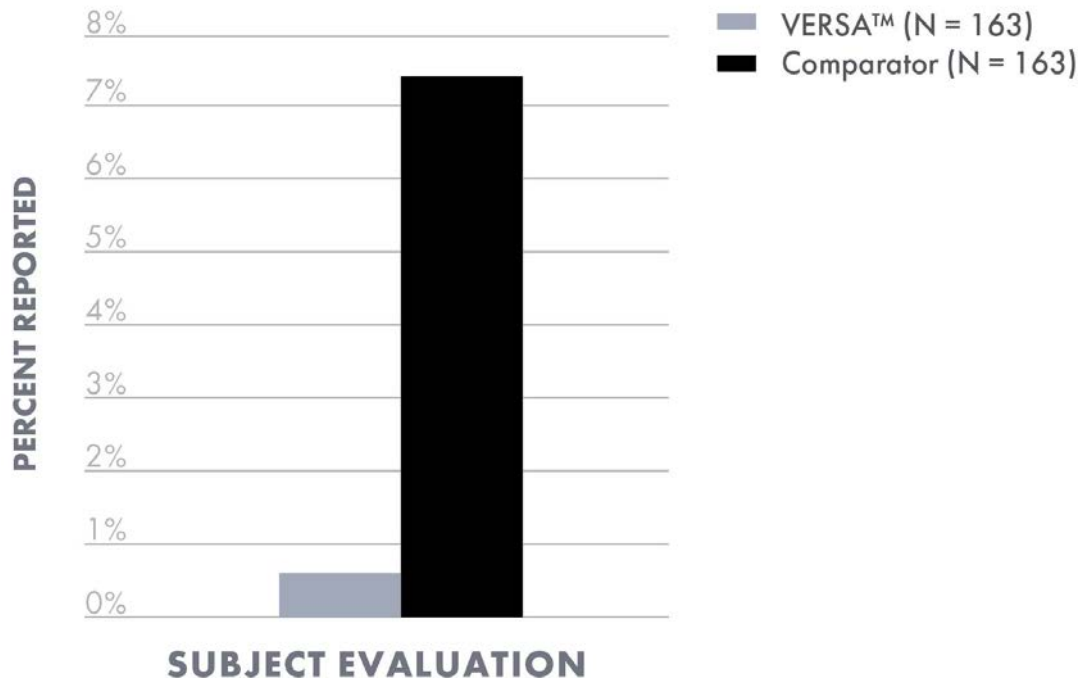
Revanesse® Versa™ Clinical Study Report SYM 2014-02: A Multicenter, Double-Blind, Randomized, Split-Face Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ versus Restylane® for the Correction of Nasolabial Folds. Revanesse® Versa™ is a registered trademark of Prollenium Medical Technologies Inc. Restylane® is a registered trademark of Nestlé Skin Health S.A.

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ITT: Intent-to-Treat subjects

# TREATMENT EMERGENT ADVERSE EVENTS

## SEVERE TEAEs AT INJECTION SITE



7.4% of the ITT subjects treated with Comparator reported severe TEAEs at the injection site vs. 0.6% of the subjects treated with Versa™

Revanesse® Versa™ Clinical Study Report SYM 2014-02: A Multicenter, Double-Blind, Randomized, Split-Face Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ versus Restylane® for the Correction of Nasolabial Folds. Revanesse® Versa™ is a registered trademark of Prollenium Medical Technologies Inc. Restylane® is a registered trademark of Nestlé Skin Health S.A.

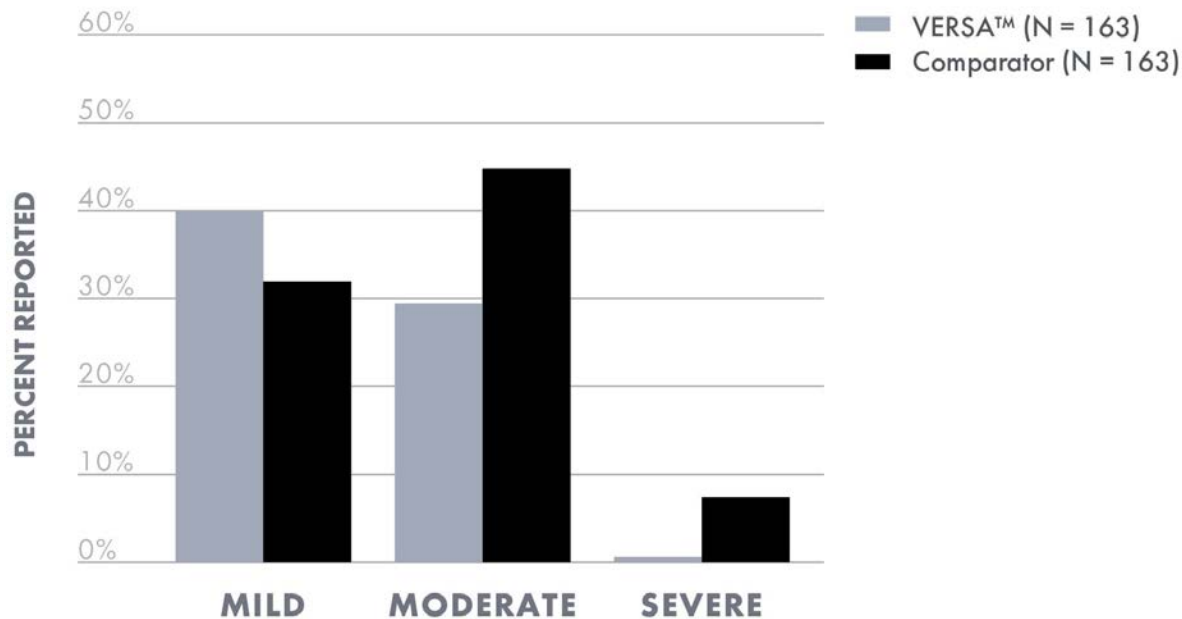
For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)

ITT: Intent To Treat subjects



# TREATMENT EMERGENT ADVERSE EVENTS

## SUMMARY OF INJECTION SITE TEAEs



84.0% of the ITT subjects treated with Comparator reported TEAEs vs. 69.9% of the subjects treated with Versa™

Revanesse® Versa™ Clinical Study Report SYM 2014-02: A Multicenter, Double-Blind, Randomized, Split-Face Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ versus Restylane® for the Correction of Nasolabial Folds. Revanesse® Versa™ is a registered trademark of Prollenium Medical Technologies Inc. Restylane® is a registered trademark of Nestlé Skin Health S.A.

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ITT: Intent To Treat subjects

# CLINICAL TRIALS HIGHLIGHTS

**REVANESSE® VERSA™ HAS BEEN PART OF THREE CLINICAL TRIALS, INVOLVING MORE THAN 300 PATIENTS.**

- Multicenter Double-Blind Randomized Split-Face Study to Evaluate Revanesse® Versa™ vs. Restylane® for the Correction of Nasolabial Folds
- Multicenter Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ Retreatment
- Study to Evaluate the Safety and Efficacy of Revanesse® Versa™ + vs. Revanesse® Versa™ for the Correction of Nasolabial Folds

## STUDIES DEMONSTRATE

- Over 300 subjects treated with no serious adverse events reports
- Non-inferiority to Restylane
- Less swelling
- 1-year pGAI data with optimal correction (subjects did not go back to baseline)

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**REAL PEOPLE  
REAL RESULTS**



# REAL PEOPLE REAL RESULTS

## BEFORE & AFTER

Nasolabial folds treated with Versa™.  
Results may vary for other patients.



REVANESSE® VERSA™

✦ Made in Canada

# REAL PEOPLE REAL RESULTS

## BEFORE & AFTER

Nasolabial folds treated with Versa™.  
Results may vary for other patients.



REVANESSE® VERSA™

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# REAL PEOPLE REAL RESULTS

## BEFORE & AFTER

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# REAL PEOPLE REAL RESULTS

## BEFORE & AFTER

Nasolabial folds treated with Versa™.  
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REVANESSE® VERSA™

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# REAL PEOPLE REAL RESULTS

## BEFORE & AFTER

Nasolabial folds treated with Versa™.  
Results may vary for other patients.



BEFORE



AFTER

REVANESSE® VERSA™

✦ Made in Canada



# REAL PEOPLE REAL RESULTS

## BEFORE & AFTER

Nasolabial folds treated with Versa™.  
Results may vary for other patients.



BEFORE



AFTER

REVANESSE® VERSA™

✦ Made in Canada

# REAL PEOPLE REAL RESULTS

## BEFORE & AFTER

Nasolabial folds treated with Versa™.  
Results may vary for other patients.



BEFORE



AFTER

REVANESSE® VERSA™

✦ Made in Canada

The background of the entire page is a light gray with several clear, realistic water droplets of various sizes. Some droplets are in sharp focus, showing highlights and reflections, while others are blurred in the background, creating a sense of depth. The droplets are scattered across the right side and bottom of the page.

# PROLLENIUM®

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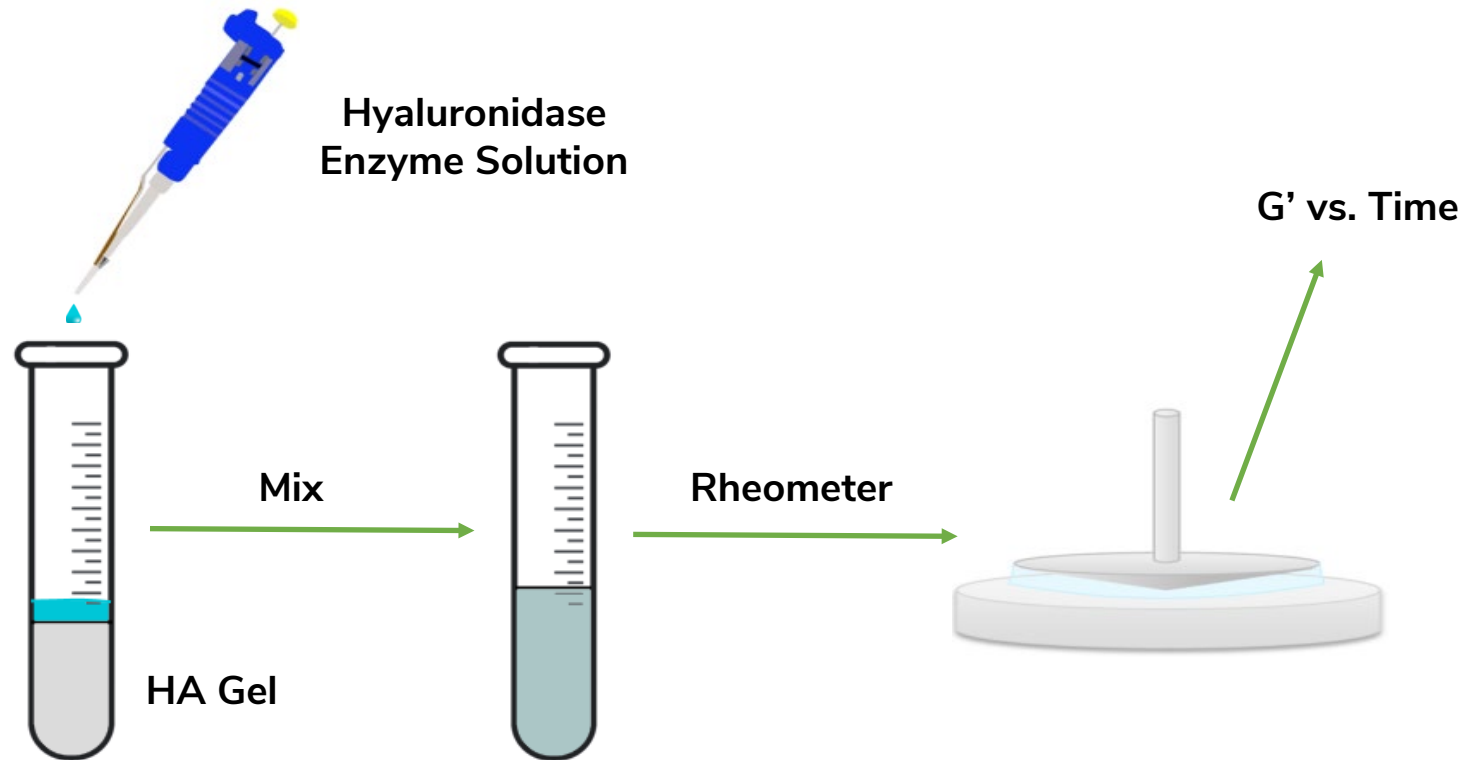
[RevanesseUSA.com](http://RevanesseUSA.com)





# APPENDIX

# DEGREDDATION TESTING



For more information on Revanesse® and Important Safety Information, please visit [RevanesseUSA.com](http://RevanesseUSA.com)