

Disclosures

- I have received honoraria in the past 2 years for speaking, writing, participating in an advisory capacity, research or meeting support from: Apellis, ABB Optical, Alcon Laboratories, Aldeyra, Allergan, Art Optical, Avellino, Bausch + Lomb Health, Contamac, Cooper Vision, CSEye, Dompe, Horizon Therapeutics, Johnson & Johnson Vision Care, Kala, Lenstech, Notal Vision, Novartis, Oyster Point, Quidel, Radius XR, RVL, Sun Pharma, Tangible Science, Tarsus, Santen, Visus, Visionix, Walman Optical and Zea Vision.
- All relevant relationships have been mitigated



InflammaDry

- Detects elevated levels of MMP-9 in tear fluid
- Rapid 10 minute results
- Easy to use can be performed by a nurse or technician
- In-office (point of care) test
- Low cost no additional equipment required
- Low cost no additional ed
 One time use disposable
- Accurate high sensitivity and specificity



 $InflammaDry\ is\ CE\ Marked\ and\ commercially\ available\ in\ Europe.\ At\ this\ time\ InflammaDry\ is\ pending\ 510(k)\ review\ by\ FDA\ and\ is\ not\ commercially\ available\ in\ the\ U.S.$

How to Use InflammaDry: Four-step Process



- Gently dab the Sample Collector in 6-8 locations on the palpebral conjunctiva (lower eyelid) to collect a tear sample. Do not use a dragging motion.
- . Snap the sample collector into the test cassette and press firmly where indicated.
- Dip the test cassette into the provided buffer vial for 20 seconds. Replace the cap.
- 4. Read the results: 2 lines (1 red, 1 blue) = positive, 1 line (blue) =







Signal Strength

Osmolarity Digital Readout



Tear-Based Point-of-Care (T-POC) Quantitative Testing Platform

- . Used to improve patient care in the areas of:
 - Dry eye disease (DED)
 - Ocular immunology,
 - · Ocular allergy,
 - Ocular surgery, including LASIK
- With T-POC in your practice, patients can be diagnosed, treated, and monitored directly within your practice
 - Quantitative IgE and Lactoferrin results delivered within minutes
- Just a 1 microliter tear sample from each eye required
- Billable & reimbursable by Medicare and Private Insurance*
 - CPT 83520 (Lactoferrin)
 - CPT 82785 (IgE)

T-POC Lactoferrin Testing Is it Aqueous Deficient or Evaporative Disease?

- Benefits of testing Lactoferrin levels in the tear film:
 - Low Lactoferrin levels (<0.9 ug/ml) directly correlates to DED caused by aqueous deficiency
 - Severity of DED can be determined by the Lactoferrin level
 - Lactoferin level

 Low Lactoferin levels indicate DED and depressed ocular immunity, which may represent an increased surgical risk

 Low Lactoferin levels may indicate the cause of contact lens intolerance

 Changes in Lactoferin levels may show the efficacy of the prescribed treatment

 Dry eye is a relative contraindication for LASIK and other surgical procedures



T-POC IgE Testing: Is There An Allergic Component?

- Benefits of testing IgE levels in the tear film:
 Presence of IgE indicates the diagnosis of allergic conjunctivitis
 Levels of IgE increase with the severity of the allergic response
 IgE testing can help differentiate allergic conjunctivitis from DED
- Conjunctivitis from DED

 Changes in IgE levels may show the efficacy
 of prescribed treatment
 IgE value is < 80 ng/mL (33 kIU), there is a
 95.7% probability that the patient does not
 have an ocular allergy
- IgE value is > 80 ng/mL, there is a 92.9% probability that this elevated IgE is indicative of an ocular allergy



Tear-Based POC Quantitative Testing









No AMD -No drusen or <10 small drusen (<63um) without pigment abnormalities Early AMD -Approximately >9 small drusen or <15 intermediate drusen (63-124um) or pigment abnormalities associated with AMD Intermediate AMD -Approximately >14 intermediate drusen or any large drusen (>124um) Advanced AMD -Geographic atrophy with involvement of the macular center or non-central GA atleast 350um in size



Real-World Remote Patient Monitoring Model

Preferential Hyperaculty Perimetry test for detecting visual abnormalities

Patient's Home

Preferential Hyperaculty Perimetry (PHP) identifies visual distortions caused by, but not limited to, conversion from dry to wet AMD

Foreseet-from (Notal Vision) uses PHP test on a home-based device performed by the patient

DID YOU KNOW?

-1M people are affected by GA in the US

ision loss from GA can occur before tesions reach the fovea

The median time to foveal encroachment in GA patients was just 2.5 years

SYFOVRE APPROVED ON FEB 17, 2023

INDICATION: SYFOVRE™ (pegcetacoplan injection) is indicated for the treatment of geographic atrophy (GA) secondary to agerelated macular degeneration (AMD).

INTRAVITREAL INJECTION given monthly or every other month (EoM); Based upon doctor's discretion

[17]

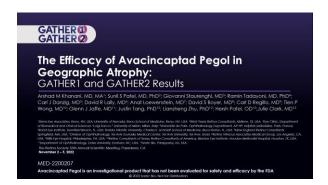
Mechanism of Action: Targeting C3 within the Complement Cascade

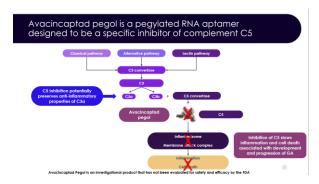


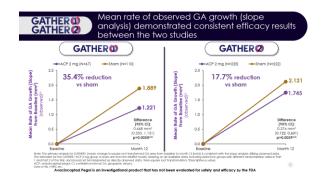
RESULTS: Monthly and EoM











Avacincaptad pegol (IZERVAY) is approved for the treatment of geographic atrophy secondary to age-related macular



- Efficacy and safety of IZERVAY were evaluated in 2 phase 3, randomized, multi-center, doublemasked, sham-controlled clinical trials
 - Izervay safety profile was consistent, with a high rate of patient retention
 - Izervay demonstrated significant reductions in ga growth in just 12 months

^{1.}JZERVAY™ (av acincaptad pegal intravitreal solution) Prescribing Information, IVERIC bio, Inc., Parsippany, N

New Data on Zeaxanthin

Prevention Surpasses Treatment: 5-year Follow-Up, Cost-Utility, and Cost-Benefit of Zeaxanthin Therapy for Neovascular Age-Related Macular Degeneration

Received: March 20, 2023 / Accepted: May 22, 2023 © The Author(s) 2023

ABSTRACT

-Group randomized to 20mg of Zeaxanthin (ZX) per day + AREDS 2 vs just AREDS 2

-Paitents developing nAMD in the nonaffected eye in that time period

-22% of ZX -48% of non-ZX



iCare HOME2 and HOME

Available measurement positions The new iCare HOME2 enables measurement freedom

iCare HOME2

iCare HOME

Measurements are possible in sitting position only



iCare HOME2 and HOME

Measurement guidance before the measurement

Smart light guide enables active positioning guidance for the patient. The light rings help find the correct measurement distance.

- Solid red light means the device is not level and cannot take a
- Visible **blue light ring** indicates that the probe is too far from the eye
- When the patient sees a
- symmetrical green light ring, the device is positioned in correct angle

- Probe base light to help level the tonometer
 - Solid red light means the device is not level and cannot take a measurement
 - When the patient sees a symmetrical green light ring, the device is level and centered

iCare HOME2 and HOME Taking measurements

- check mark, shown on the display screen and single long beep
- pressing Measure button 6 individual times or
- pressing and holding the Measure button down

- pressing Measure button 6 individual times or



Wearable Devices for VF

Radiusxr Olleyes Heru

M&S Technologies











Study on Effect of Nutrition on Computer Symptoms

- 360 participants
- Randomized to 4 groups that utilized 1 of four nutritional formulations for 90 days
- Measurements were taken at baseline, 45 days later and 90 days later
 - 1 Placebo

 - 1 Haceou
 2 6 mg Lutein, 1.2mg Zeaxanthin, 75mg chrysanthemum extract, 75mg goji berry extract, 100mg black current extract
 3 10 mg Lutein, 2 mg Zeaxanthin, 125mg chrysanthemum extract, 125mg goji berry extract, 167mg black current extract
 4 14 mg Lutein, 2.8 mg Zeaxanthin, 175mg chrysanthemum extract, 175mg goji berry extract, 233mg black current extract

Results

- 1 Improved eye fatigue symptoms
- 2 Improved visuognosis persistence
- 3 Improved Schirmer test
- 4 Improved macular pigment optical density (MPOD)

2. Improved visuognosis persistence

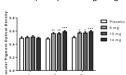
- Subject stares at an object for 3 minutes
- · As subject stares at object, it will at some point blur
- The percentage "clear vision" time of the total gaze time is called visuognosis persistence, which is an index that reflects both visual and psychological functions
- After staring at the cubes for a while, the subjects would see inverted cubes, which was defined as "unclear vision."
- Visuognosis persistence=(clear vision time/total time of gaze)×100%
- · All formulas improved visuognosis persistence

3. Improved Schirmer test

- Unanesthetized measurement over 5 minutes for each eye
- Measured the number of patients in each group with >9mm
- The highest concentration nutrition test groups showed significant improvement after 90 days over placebo

4. Improved Macular Pigment Optical Density (MPOD)

- All baseline measurements for all groups were identical
- All treatment groups demonstrated significant increase in MPOD over placebo at second visit (45 days after beginning nutrition)



Summary

- Nutritional supplementation improves eye fatigue symptoms, visuognosis persistence, Schirmer strip test and MPOD levels
- Some of symptom and sign improvement likely secondary to MPOD
- MPOD has traditionally been considered a biomarker for risk factors associated with long term disease development and progression
- MPOD may be an appropriate functional marker for eye fatigue
- Understanding the massive amount of screen time currently being performed by everyone, this is becoming an increasingly important area for eye care providers to intervene when appropriate



ELM – Eye Lipid Mobilizer (Not Yet Available)





https://www.eyedetec.com/

AZR-MD-001

- · Mechanism of action: Contains selenium sulfide, a keratolytic agent also found in some anti-dandruff shampoos.
- Safety/efficacy:
 - An interim data analysis from a double-masked, vehicle-controlled trial presented at ARVO 2021 found a significantly greater change in OSDI score of at least 4.5 units in the group receiving 1.0% AZR concentration (85.7%) vs vehicle group (22.2%, *P*=.03)

 Change from baseline in Meibomian Gland Score was also
 - significantly higher with 1.0% AZR vs control (P=.03)

Gupta PK et al. Clin Cjohthalmol. 2021;15:4399-4404. https://lone.arvojoumals.org/article.aapp https://clinicaltrials.gov/ct2/show/NCT008520517lerms.azv-md-0018.dosw-28canku-4

NOV03 (Currently Available - Miebo)

Mechanism of action: Sole ingredient is perfluorohexyloctane, which covers the tear film to limit aqueous evaporation and penetrates meibomian glands, reportedly dissolving altered meibum

Safety/efficacy:

- Safety/efficacy:

 Results of the phase 3 double-masked GOBI study presented at the 2022 ASCRS annual meeting showed that adults with MGD-related DED in the treatment group vs placebo group had significantly improved total corneal fluorescein staining (2.0 vs -1.0) and visual analog scale dryness score (2-74 vs -19.7), Po.001 for both) on Day 57.

 Blurred vision was the only adverse event more likely in treated subjects (3.0% vs 0.204).
- Topline data of the phase 3 MOJAVE trial announced in 2021 also included significant change on these efficacy outcomes versus control on day 57 (P<.001 for both)

Meiverter













Acta Ophthalmologica Acta Ophthalmologica Review Article Review of the impact of presbyopia on quality of life in the developing, and developed world Article Review of the impact of presbyopia on quality of life in the developing, and developed world Article Article

Current Treatments (Non-surgical)

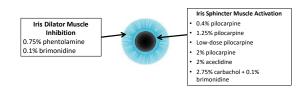
- Glasses
 - Reading glasses
 - Advantages low cost, easily accessible, convenient
 - Disadvantages inconvenient, are dependent on them
 - Bifocals
 - Advantages have correction on all the time (both distance and near)
 - Disadvantages sometimes current environment isn't conducive to this design; cosmetic component
 - Progressive addition glasses
 - Advantages have correction on all the time (both distance and near)
 - Disadvantages sometimes current environment isn't conducive to this design

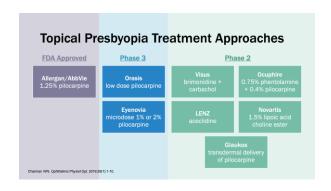
Current Treatments (Non-surgical)

Contact Lenses

- Single Vision Distance
 - Advantages provides visual freedom from spectacles
 - Disadvantages need optical correction for near vision
- Monovision
 - Advantages provides visual freedom from spectacles
 - Disadvantages some visual compromise
- Multifocals
 - Advantages provides visual freedom from spectacles
- Disadvantages some cannot adapt to multifocal optics

MOAs of Different Miotics In Development For Presbyopia







Ocular Surface Inflammation

- Commercially available as Restasis
 - 0.05% concentration
 - Preservative free
 - Single vials
 - Multi-dose bottle
 - Patented non-preserved system that maintains its sterility
- Is an immunosuppressive agent
- Available as generic

Restasis (Cyclosporine 0.05%)

Cyclosporine 0.09%

- •Commercially available as Cequa
- Unit dose vials
- •Unique delivery mechanism
- N-cell technology

Klarity-C Drops PF

- 0.1% cyclosporine
- Preservative free
- Imprimis Rx
- 5.5mL multidose bottle



Verkazia

- 0.1% cyclosporine
- Approved for vernal keratoconjunctivitis
- Dosed four times/day



Cyclosporine 0.1%

- Novaliq
- VEVYE
- FDA approved for the signs and symptoms of dry eye
- Water free

Ocular Surface Inflammation

- Intercellular adhesion molecule-1 (ICAM-1) is found on surface cells of the conjunctiva and cornea
- ICAM-1 is overexpressed in dry eye disease
- •LFA-1 and ICAM-1 binding is believed to be propagate inflammatory cascade on the ocular surface
 - Is believed to do so by T-cell activation and migration to target tissues

Ocular Surface Inflammation

- Lifitegrast applied topically is believed to prevent the interaction between LFA-1 on T-cells and ICAM-1 on the ocular surface
- Has been shown to improve both signs and symptoms of dry eye within a 12 week time period
- · Commercially available as Xiidra
 - 5% concentration
 - non-preserved sterile unit dose vials
- It is a lymphocyte functioning antigen-1 (LFA-1) antagonist

Tyrvaya



RASP Inhibitors

- · www.aldeyra.com
- Aldehyde is a product of metabolism
- Normally rapidly broken down
- With inflammation, is produced in quantities that are difficult to breakdown efficiently
- RASP Reactive aldehyde species
- Reproxalap Is a RASP inhibitor



Effectiveness of a Singular Ocular Rinse via Irrigating Eyelid Retractor to Reduce MMP-9 in Patients with Dry Eye Disease

Nandini Venkateswaran, MD ABO Natasha Mayer, BSc

ASCRS 2023

San Diego, California

Objective

Eye rinsing has been an effective method to reduce Matrix Metalloproteinase-9 (MMP-9), a hallmark of surface inflammation

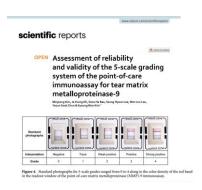
A single ocular rinse assisted via irrigating eyelid retractor effect on MMP-9

Irrigating Eyelid Retractor

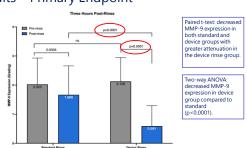
Fixed to a syringe, the retractor has 5 ports which aim fluid at the palpebral conjunctiva, bulbar conjunctiva and conjunctival fornix.







Results—Primary Endpoint



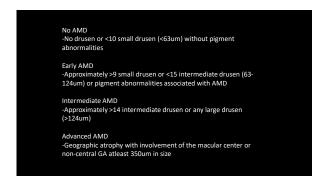
Conclusion

A novel irrigating eyelid retractor rinse of the ocular surface statistically reduces MMP-9 levels compared to baseline and is superior to a standard eye rinse.

Use of an irrigating eyelid retractor may be a therapeutic avenue for those patients with dry eye disease.

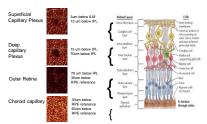
Further work on the durability of these findings is ongoing.







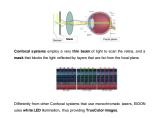
Visualize Retinal Vasculature





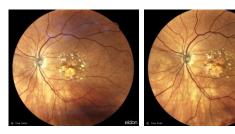


TrueColor Confocality

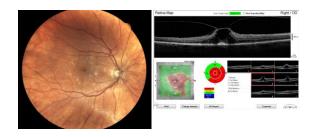




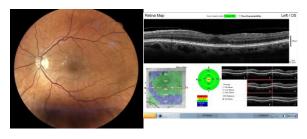
New Standards for Detecting Change



Macular findings



Macular findings









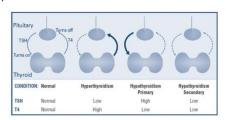




Thyroid Function

- TSH is produced by pituitary gland
- \bullet Thyroid Gland Produces T4 which is converted to T3
- Alterations in hormonal balance can cause problems to arise

Thyroid Gland Function

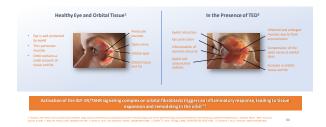


https://www.thyroid.org/thyroid-function-tests/

TED Is a Debilitating, Progressive, and Vision-threatening Autoimmune Disease



Inflammation, Tissue Expansion, and Eye Muscle Changes May Lead to the Clinical Manifestations of TED



Recognizing the Signs and Symptoms of TED

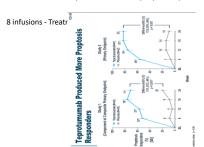


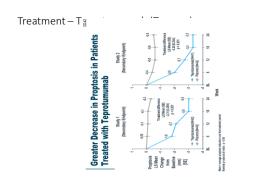
Ongoing Inflammation and Expansion of Orbital Tissues Leads to Changes in Physical Appearance



Billion ED, et al. Clin Ophishimal 2009, INSENDE S. E. Bahn NS. Et algo / Ether 2010, NEXT 2011, E. Eddor C, et al. Epethanica. Op. 1034 / Performance and independent Anticipation of participation of participation of the Section of Ministry (Section 2011). The Computer Section of Ministry (Section 2011) of the Section Operation of Section Operation Operation of Section Operation Operation of Section Operation Operation of Section Operation Oper

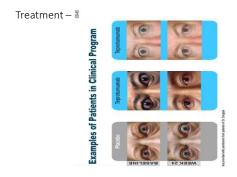
Treatment – Teprotumumab (Tepezza)





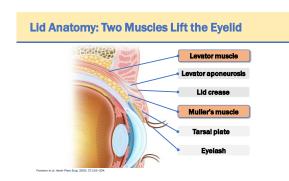
Teprotumumab Improved Diplopia

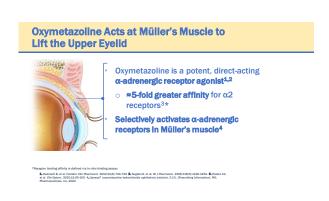
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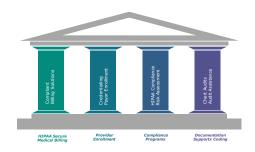












HOW IT WORKS





TE#M