






Next-Generation Therapies Transforming Patient Care Today

April 2015

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Products and Programs



Program	Targeted Procedure/ Disease	Pre-clinical	Phase 1	Phase 2	Phase 3	FDA Approval	Commercial Launch	Economic Rights
Commercial Products								
Omidria™ (OMS302) - Ophthalmology	Cataract Surgery or Intraocular Lens Replacement	<div></div>						 OMEROS
Clinical Programs								
OMS103 - Arthroscopy	Arthroscopic Meniscectomy	<div></div>						 OMEROS
MASP (OMS721) - Complement-Mediated Disorders	TMA (aHUS, TTP, Transplant-Related TMA)	<div></div>						
PDE10 (OMS824) - CNS Disorders	Schizophrenia	<div></div>						
PDE10 (OMS824) - CNS Disorders	Huntington's Disease	<div></div>						
PPARγ (OMS405) - Addiction	Opioid and Nicotine Addiction	<div></div>						
OMS201 - Urology	Ureteroscopy	<div></div>						
Preclinical Programs								
PDE7 (OMS527) - CNS Disorders	Addictions and Compulsive Disorders; Movement Disorders	<div></div>						 OMEROS
Plasmin (OMS616) - Bleeding Disorders	Surgical and Traumatic Bleeding	<div></div>						
MASP (OMS906) - Alternative Pathway Disorders	Paroxysmal Nocturnal Hemoglobinuria	<div></div>						
GPR17 - CNS Disorders	Demyelinating Disorders	<div></div>						
GPCR Platform	Multiple Disorders Across Therapeutic Areas	<div></div>						
Antibody Platform	Multiple Disorders Across Therapeutic Areas	<div></div>						

Experienced Management with Deep Industry Experience



	Position	Background
Gregory Demopulos, MD	<i>Chairman, President & CEO</i>	Stanford and Duke Departments of Orthopedic Surgery
Tim Duffy	<i>VP, Business Development</i>	MDRNA, Prometheus, Procter & Gamble
Kenneth M. Ferguson, PhD	<i>VP, Development & CDO</i>	ICOS, Cold Spring Harbor Laboratory
George Gaitanaris, MD, PhD	<i>VP, Research & CSO</i>	Nura, Primal, NCI
Patrick Gray, PhD	<i>Scientific Fellow</i>	Accelerator, Nura, MacroGenics, ICOS, Genentech
Michael Jacobsen	<i>VP, Finance & CAO</i>	Sarepta, ZymoGenetics, ICOS, Onvia, Mosaix
Marcia Kelbon, JD	<i>VP, Patent & General Counsel and Secretary</i>	Christensen O'Connor Johnson Kindness
William J. Lambert, PhD	<i>VP, Chemistry, Manufacturing and Controls</i>	MedImmune, Pacira, Eisai, Pfizer, Upjohn
Catherine Melfi, PhD	<i>VP, Regulatory Affairs and Quality Systems</i>	Eli Lilly, Indiana University
Patricia Sandler	<i>VP, Sales & Marketing</i>	Sunovion, J&J, SmithKline Beecham, Pfizer
J. Steven Whitaker, MD, JD	<i>VP, Clinical Development & CMO</i>	Allon Therapeutics, ICOS



Omidria™ (OMS302) Ophthalmological Surgery



- Omidria™ (Phenylephrine and Ketorolac Injection) 1%/0.3%
- Approved by FDA in May 2014
- www.OMIDRIA.com

Indication

- Omidria™ is an Alpha 1-Adrenergic Receptor Agonist and Nonselective Cyclooxygenase Inhibitor Indicated for:
 - Maintaining Pupil Size by Preventing Intraoperative Miosis
 - Reducing Postoperative Ocular Pain

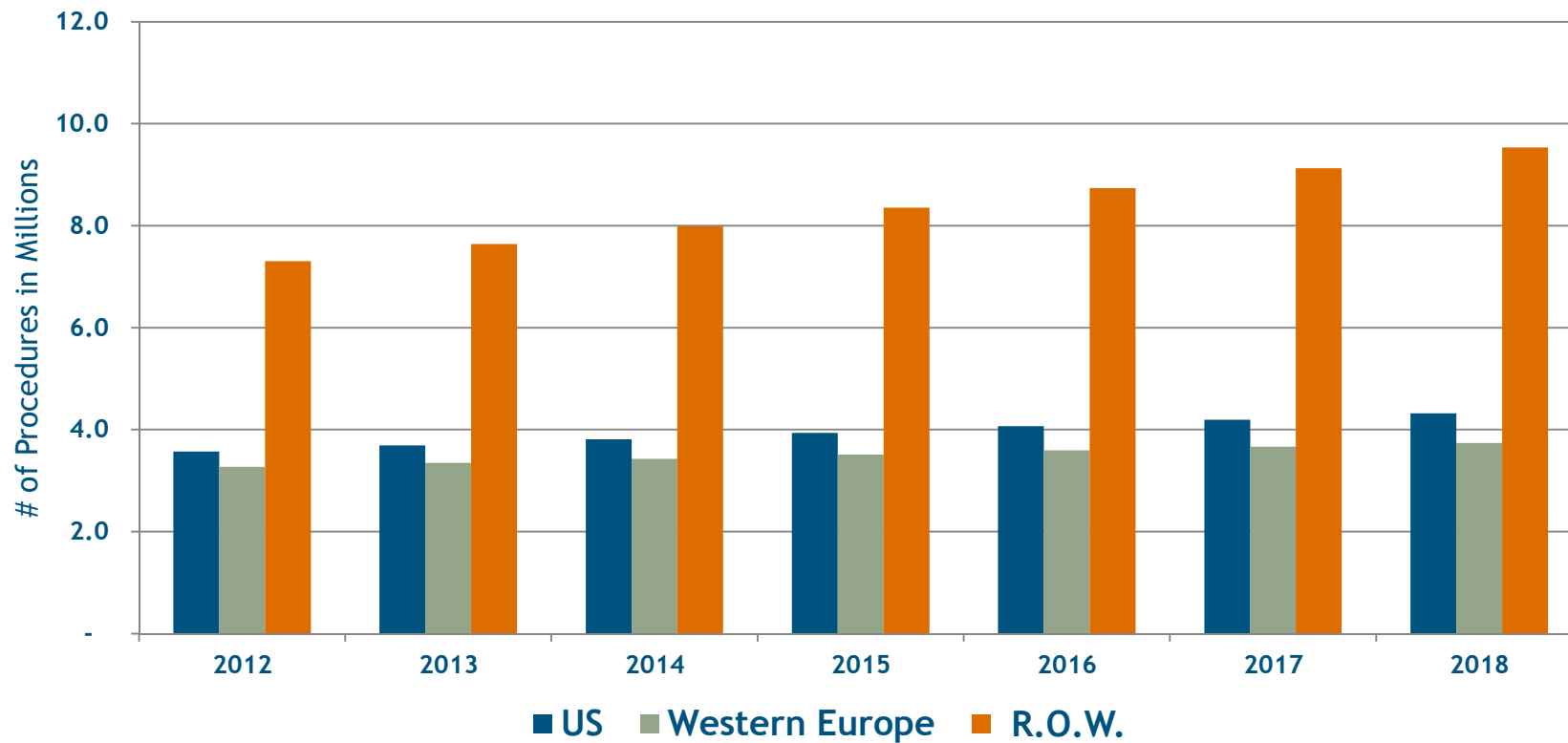
Overview

- One 4-mL Single-Use Vial Added to a 500-mL Container of Standard Irrigation Solution
- Delivers Steady-State Drug Concentrations to the Anterior Chamber throughout the Procedure
- Easily Integrated into Operating Procedural Routines
- Preservative-Free and Bisulfite-Free
- Safe and Well Tolerated - in Clinical Trials, Rates of Ocular Adverse Reactions Were Similar to Those of Placebo
- Medicare Reimbursed



IOL Replacement Surgery Market Opportunity

Lens Replacement Procedures



Source: Market Scope, Comprehensive Report on the Global IOL Market, 2011.

R.O.W. includes Japan, China, Latin America, Korea, Canada, Australia, Saudi Arabia, Taiwan, Greece, Czech, Hungary, Israel, Hong Kong, UAE, Singapore, New Zealand

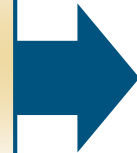


Omidria™ Fits into the Surgical Workflow



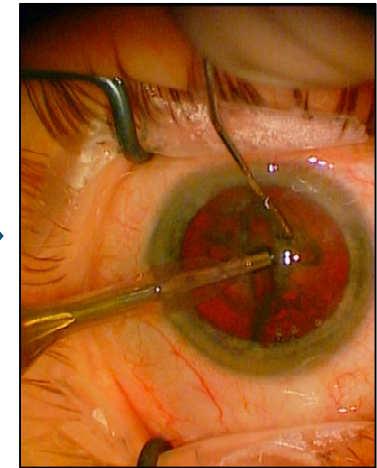
Proprietary combination of:

- Phenylephrine
- Ketorolac



4 mL

Dilute in



Once Added to Irrigation Solution, No Change in Surgical Procedure



Preoperative Topical Medication is Washed Out by Irrigation Solution

- 14 Patients Undergoing Cataract Surgery were Eligible
- Patients were Prescribed Topical Ketorolac According to the Surgeon's Usual Practice, Beginning the Day Prior to Surgery - Good Patient Compliance
- 100- μ L Aqueous Humor Samples Drawn Immediately Prior to the Initial Surgical Incision and Again Prior to Final Anterior Chamber Re-inflation and Wound Closure

Patient	Prior to Incision	Prior to Wound Closure
1	87.4	Undetectable
3	39	Undetectable
4	4.9	Undetectable
6	195	Undetectable
7	29.5	1.80
8	75.1	6.32
9	369	2.11
10	105	Undetectable
11	244	2.90
12	64.6	Undetectable
13	137	Undetectable
14	120	Undetectable

Undetectable = Below the Lower Limit of Quantification
(i.e., <1 ng/mL)

**Ketorolac Levels in the Anterior Chamber at the End of the Procedure
Were Nominal or Undetectable in All Patients**



Omidria™ Canine Study

Uptake of Ketorolac by Ocular Tissues



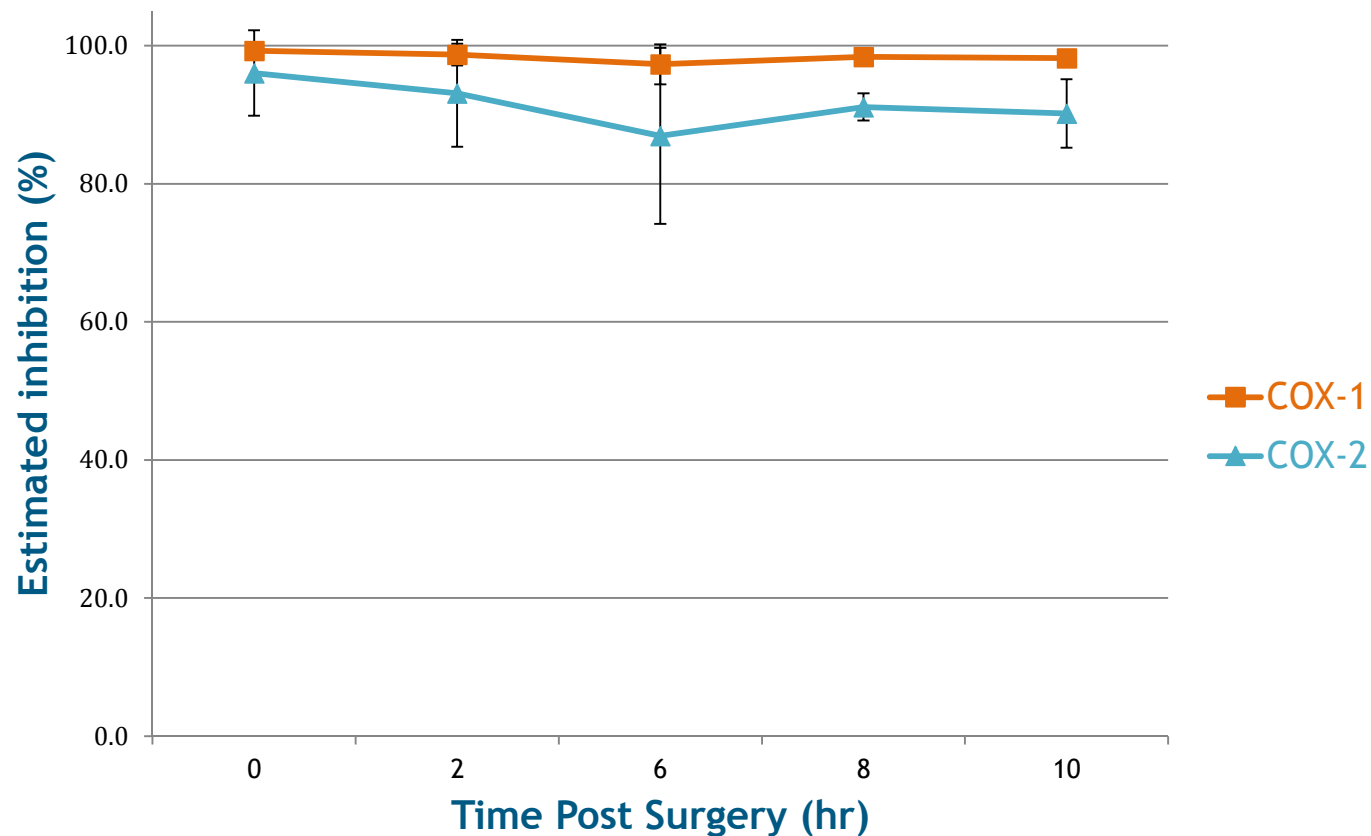
Study Design

- IOL Replacement Surgery Was Performed on 20 Canines Using OMIDRIA in Irrigation Solution
- Data Were Collected from Four Animals Per Time Point at 0, 2, 6, 8, and 10 Hours Post-procedure
- Eyes Were Enucleated and Flash Frozen; Partly Thawed Eyes Were Dissected and Retina, Choroid, Vitreous, Ciliary Body, Iris, Lens Capsule, Cornea, and Sclera Were Collected
- Ketorolac Levels Were Determined in All Samples
- Projected Inhibition of COX Enzymes Was Estimated Using IC50 Values for COX-1 and COX-2

COX = cyclooxygenase
IC = inhibitory concentration



Omidria™ Canine Study Results Estimated Retinal COX-1/-2 Inhibition



Ketorolac levels inhibited COX-1 and COX-2 Pathways from 0 to at Least 10 hours, Consistent with Phase 3 Pain Data



Omidria™ Phase 2b Trial



Summary

Trial Design

- N=221, Vehicle-Controlled, Randomized, Double-Blind Study
- Multicenter
- All Subjects Received Preoperative Mydriatics and Anesthetics
- Four-Arm, Full-Factorial Design

Co-Primary Endpoints

- Maintenance of Mydriasis (Pupil Dilation)
- Reduction of Postoperative Pain

Study Results*

- Statistically Significant and Clinically Meaningful Benefits Demonstrating Contribution of Each Component
 - *Mydriasis*
Omidria Better than:
 - Ketorolac Alone ($p < 0.0001$)
 - Vehicle ($p < 0.0001$)
 - *Pain*
Omidria Better than:
 - Phenylephrine Alone ($p = 0.0089$)
 - Vehicle ($p = 0.0418$)
- Omidria was Well Tolerated

*Presented at American Academy of Ophthalmology,
October 2011; Alan Crandall, MD



Omidria™ Phase 2b Results

Anti-Miotic Effects of Omidria and Its Components



Pupil Diameter < 6 mm				
	Vehicle* (n = 53)	Ketorolac (n = 52)	Phenylephrine (n = 49)	Omidria (n = 49)
n (%)	25 (46%)	18 (35%)	11 (22%)	3 (6%)
p-value vs. Omidria	< 0.0001	0.0004	0.0216	

*All patients, including vehicle-treated patients, received standard preoperative mydriatic and anesthetics

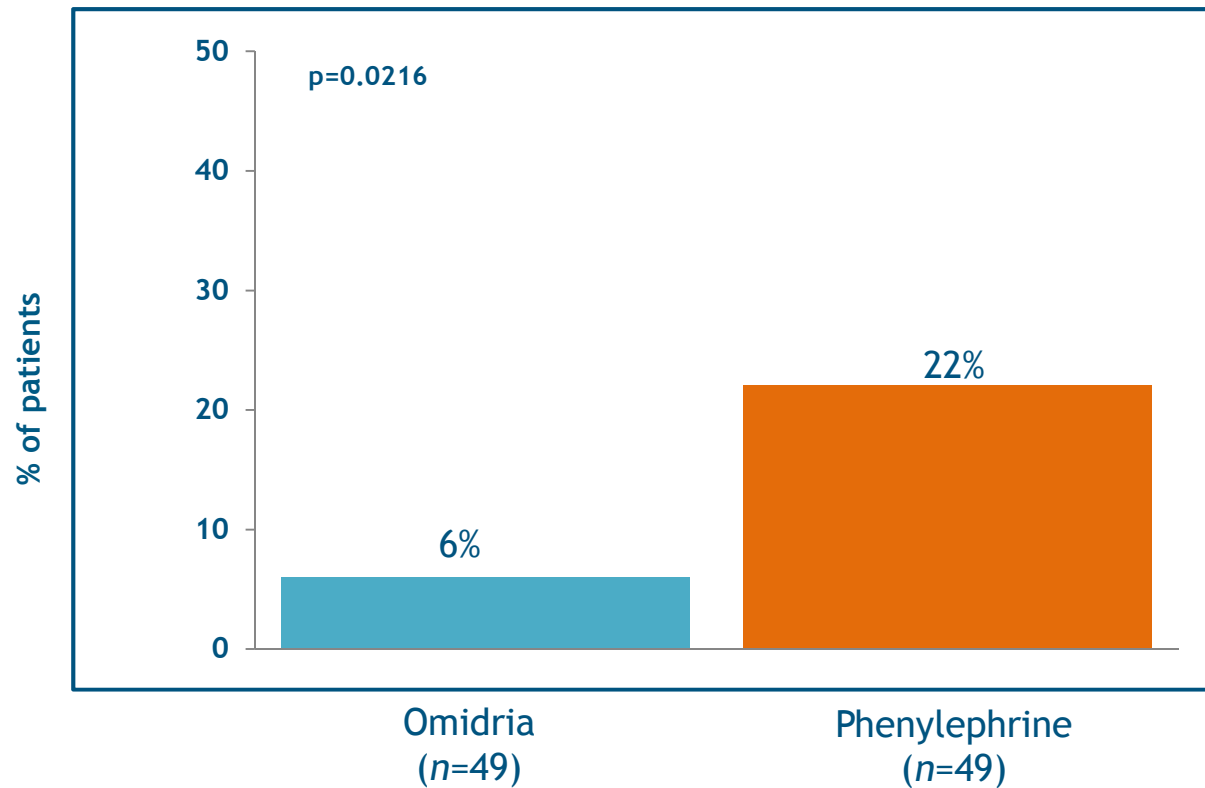
Both Ketorolac and Phenylephrine Contribute to Inhibition of Miosis



Omidria™ Phase 2b Results Omidria vs. Phenylephrine



Percent of Subjects with Pupil Diameter < 6 mm (Omidria vs. Phenylephrine)



Added to Irrigation Solution, Omidria is Superior to Phenylephrine Alone



Omidria™

Phase 3 Trials - Study 1 & Study 2



Overview

Trial Design

- Multicenter, Placebo-Controlled, Randomized, Double-Blind Study
- N=405 (ILR-003) and N=416 (ILR-004)
- All Subjects Received Preoperative Mydriatics and Anesthetics
- Two-Arm, Randomized 1:1

Co-Primary Endpoints

- Maintenance of Intraoperative Mydriasis (Pupil Dilation)
- Reduction of Pain in Early Postoperative Period*

Study Results

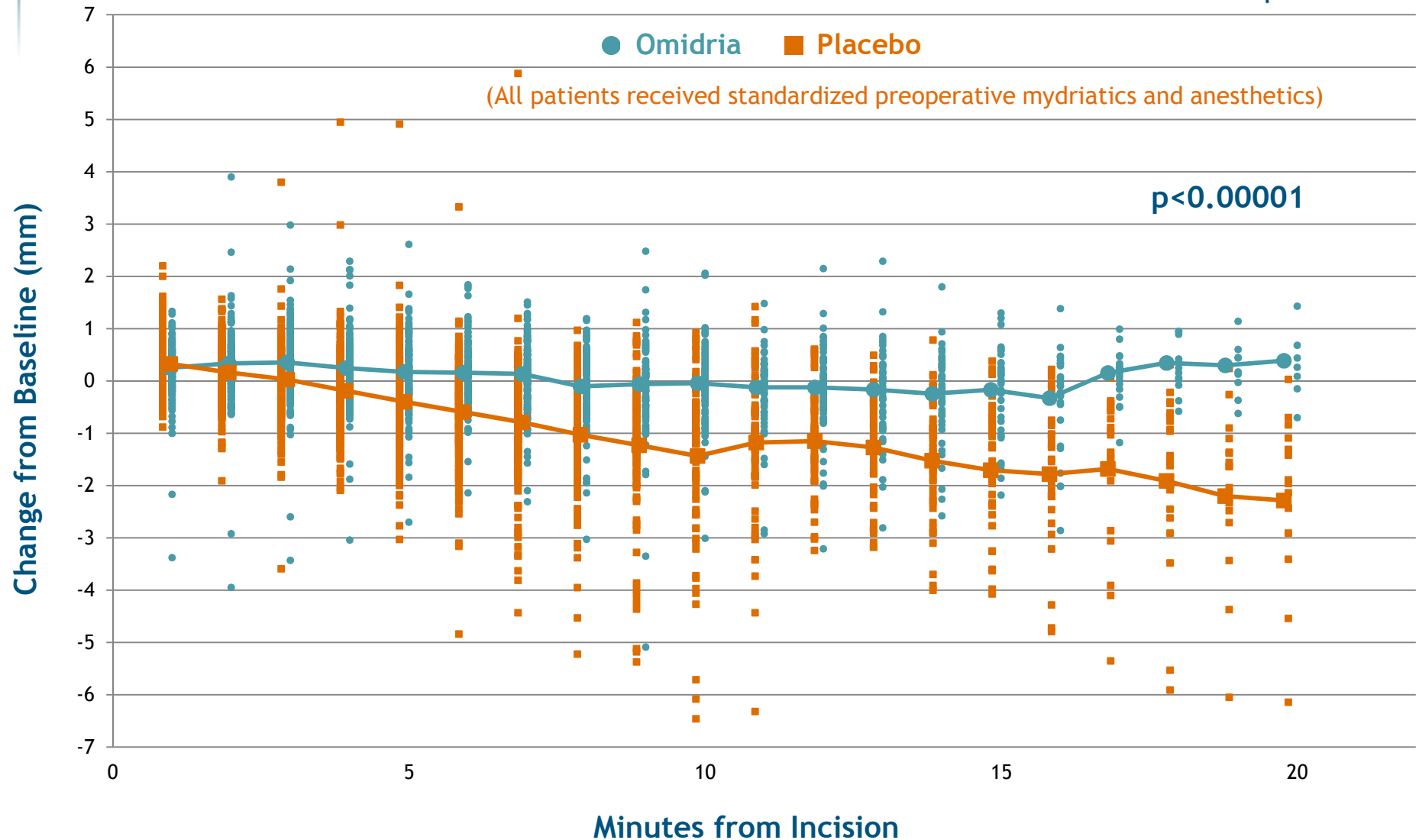
- Omidria Provided Statistically Significant and Clinically Meaningful Benefits in Both Studies
 - Mydriasis ($p < 0.00001$) in Both Studies
 - Pain ($p < 0.0001$ in ILR-003 and $p = 0.0002$ in ILR-004)
- Omidria was Well Tolerated

* Principal secondary endpoint in ILR-003



Omidria™ Phase 3 - Study 1

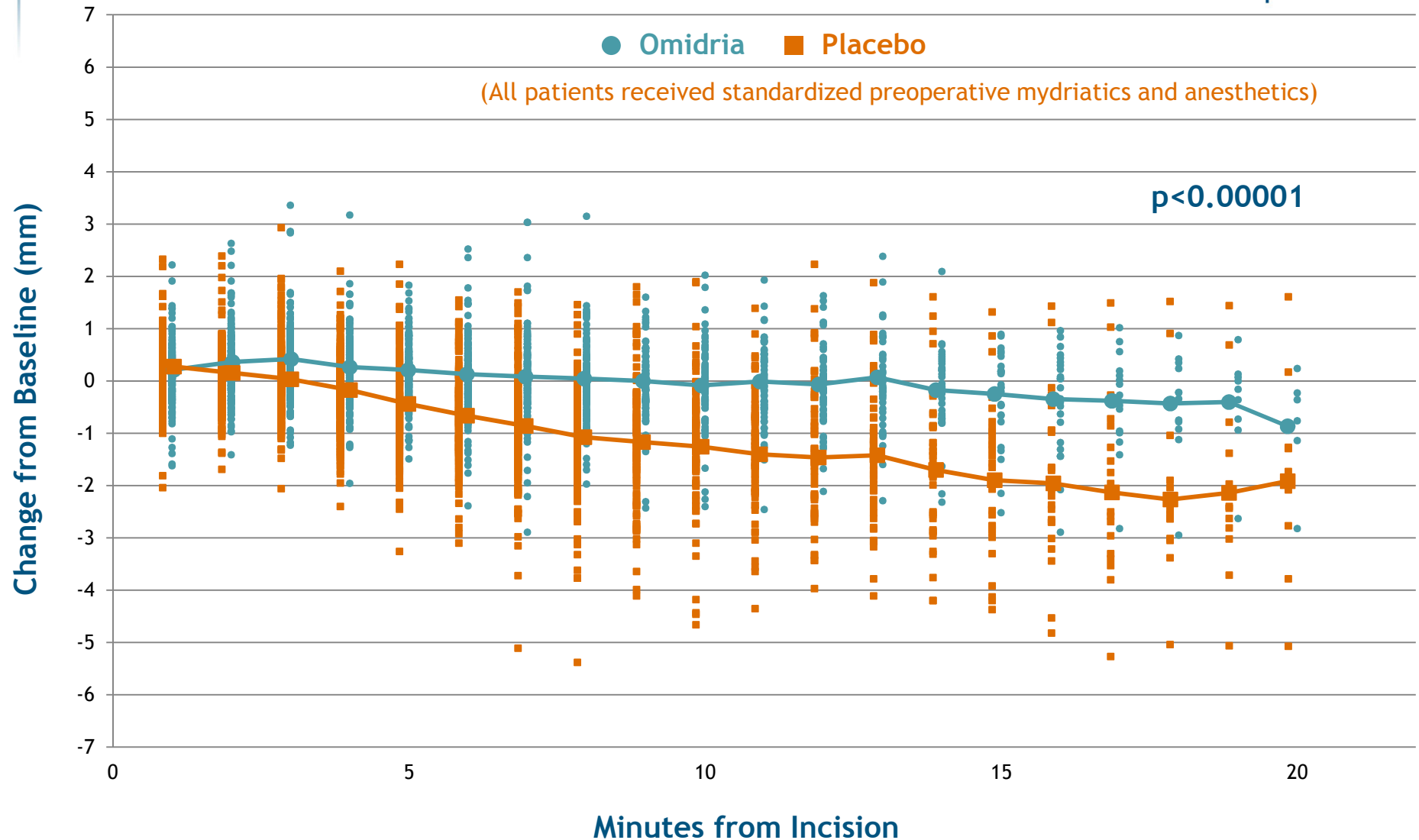
Intraoperative Change in Pupil Diameter





Omidria™ Phase 3 - Study 2

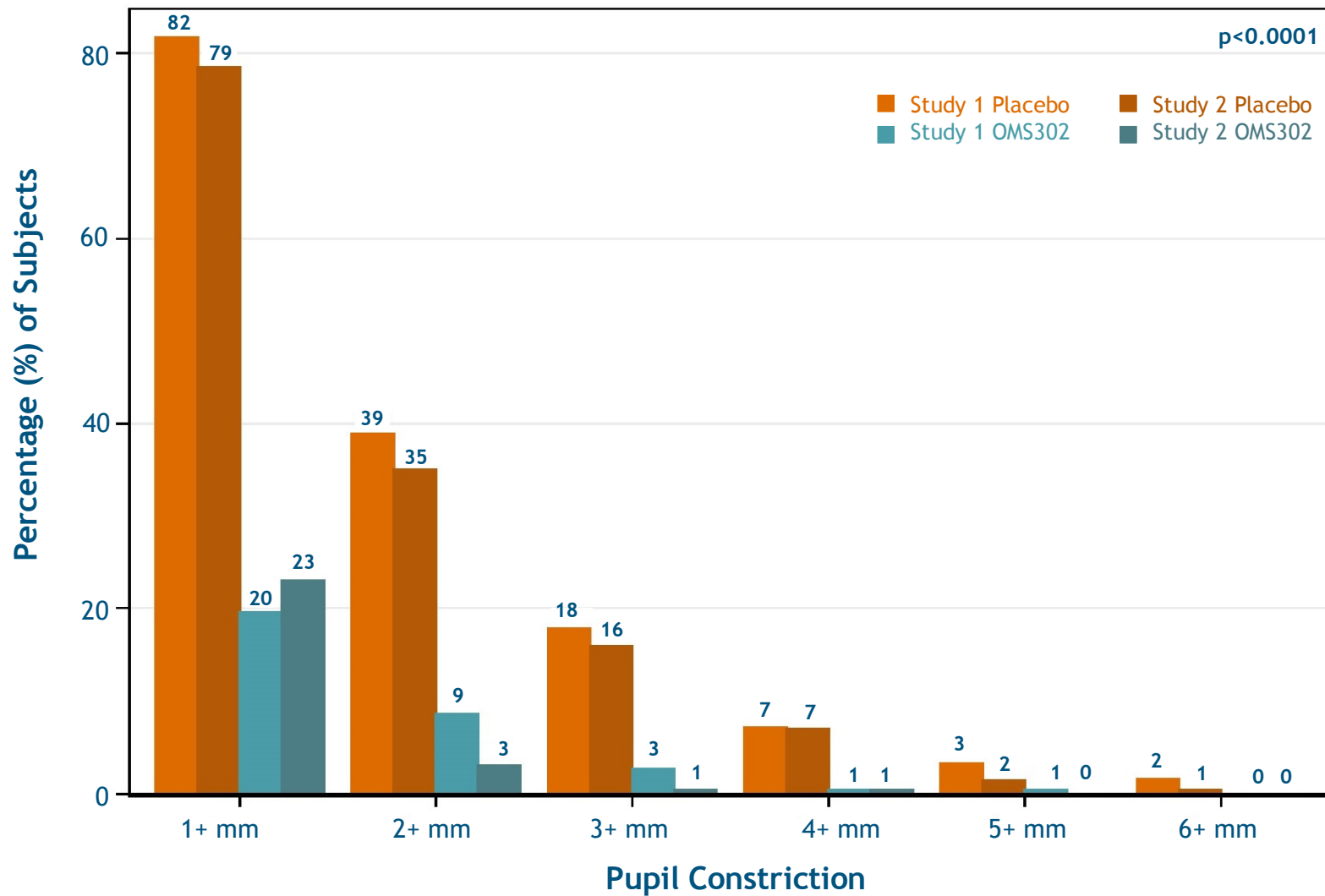
Intraoperative Change in Pupil Diameter





Omidria™ Phase 3 Trials

Studies 1 & 2- Intraoperative Pupil Constriction





Omidria™ Portal to the Operative Field

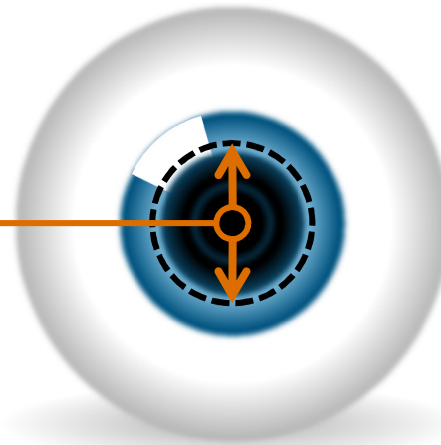


Intraoperative Pupillary Miosis

Decrease in Pupil
Diameter*
 ≥ 2.5 mm

~ 30%

Reduction in
Diameter



Resultant Decrease in
Pupil Area
(i.e., Operative Field)

~ 50%

Reduction in
Operative Field

*From a starting pupil diameter of 8.25 mm

6 mm to 3.5 mm Represents a 66% Reduction in Operative Field



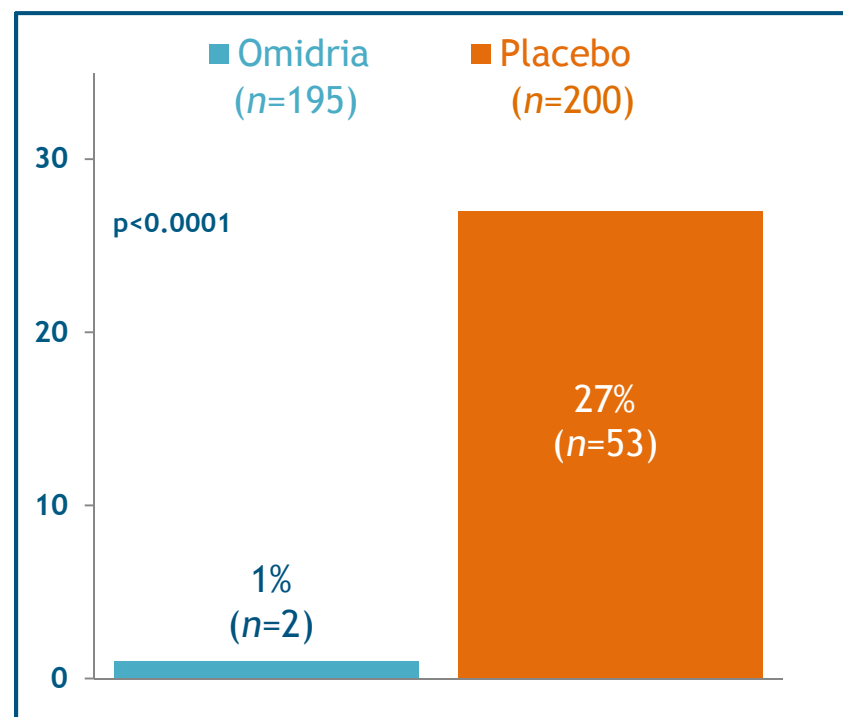
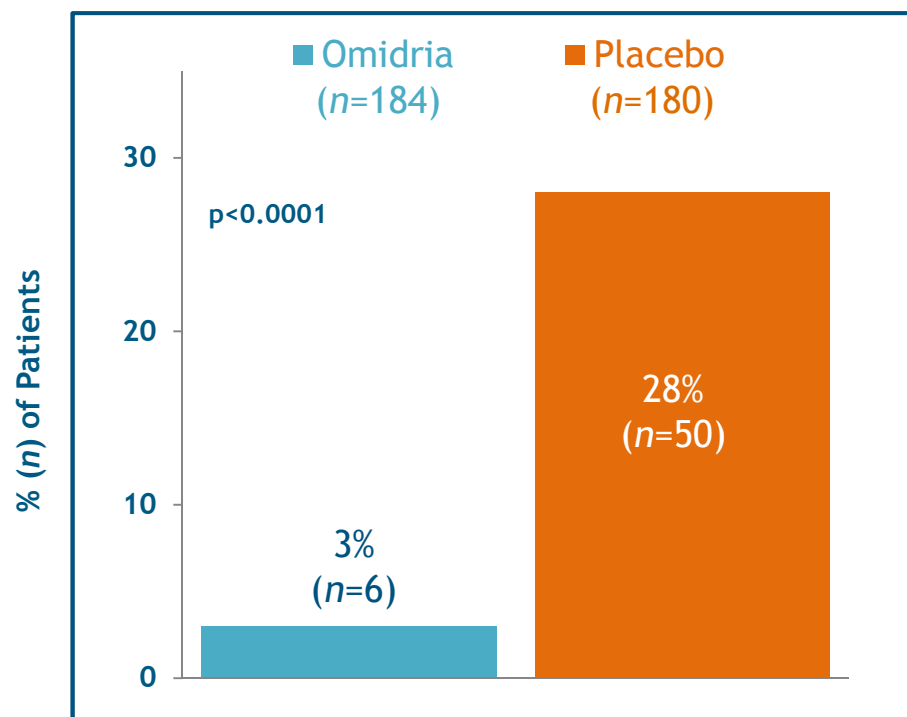
Omidria™ Phase 3 Trials

Decrease in Pupil Diameter ≥ 2.5 mm



Study 1

Study 2



*All patients, including placebo-treated patients, received standard preoperative mydriatics and anesthetics

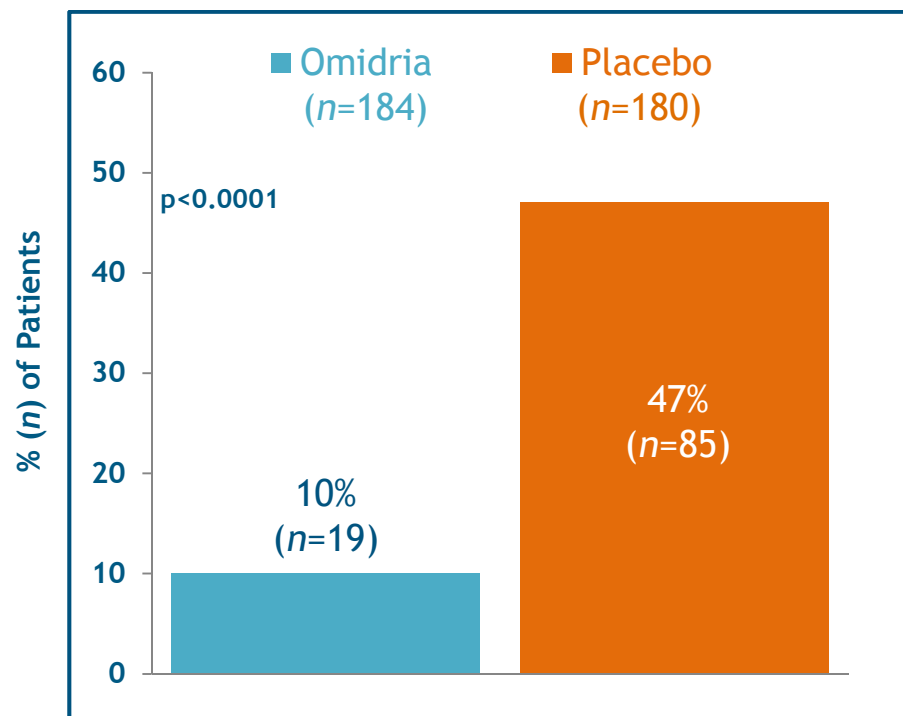


Omidria™ Phase 3 Trials

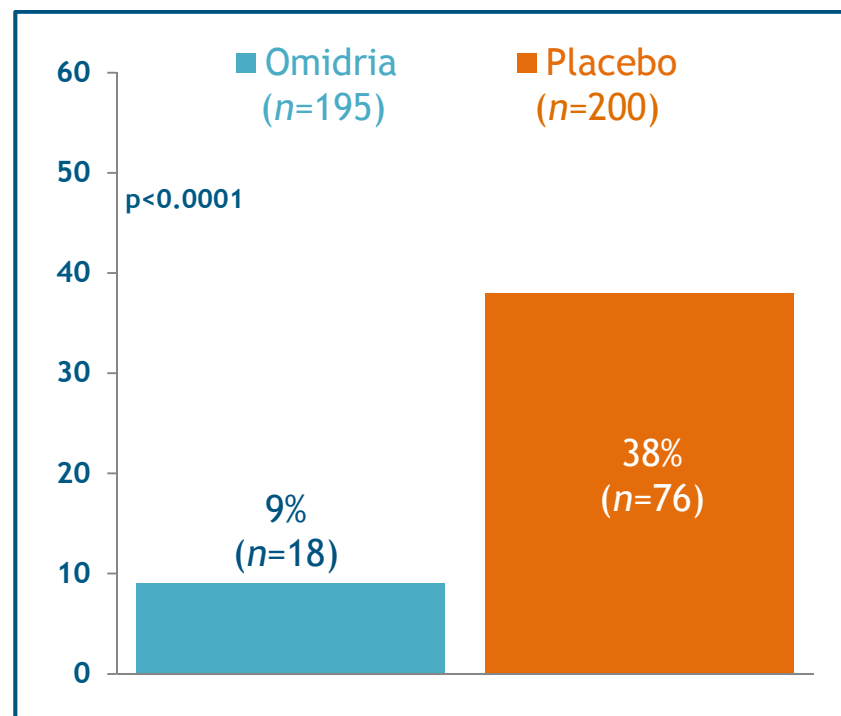
Pupil Diameter < 6 mm at Any Time During Surgery



Study 1



Study 2



*All patients, including placebo-treated patients, received standard preoperative mydriatics and anesthetics

Pupil Diameter < 6 mm is Associated with a Multiply Increased Rate of Complications (e.g., Posterior Capsule Tears, Retained Lens Fragments, and Vitreous Loss)

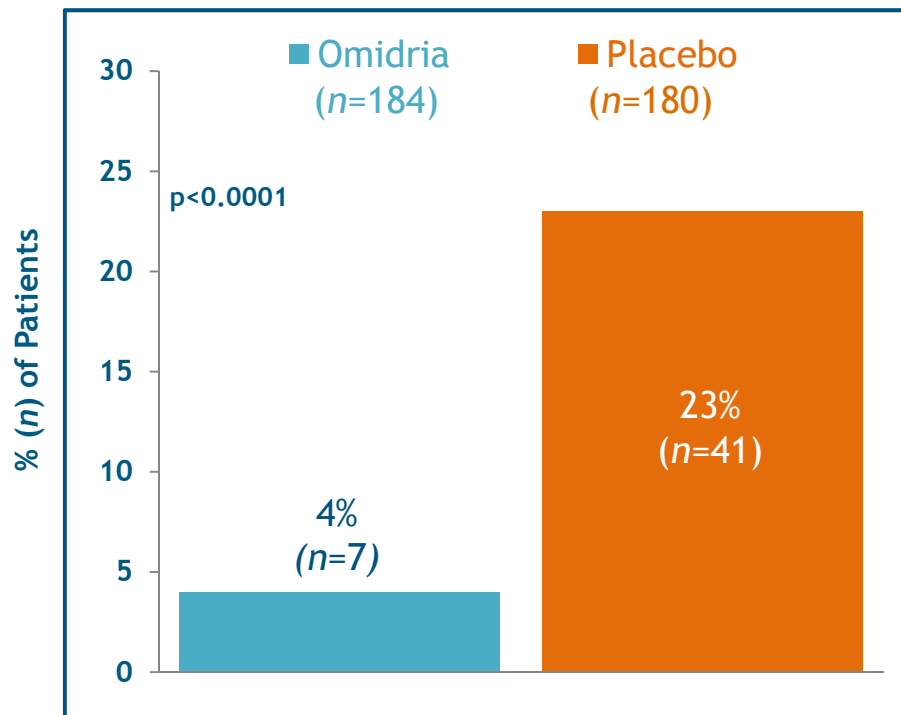


Omidria™ Phase 3 Trials

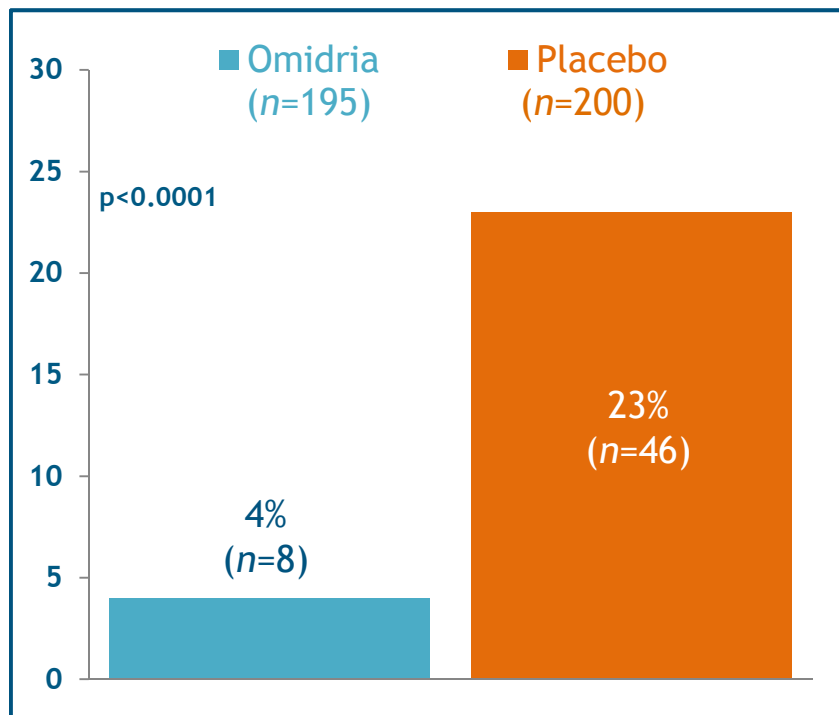
Pupil Diameter < 6 mm at Start of Lens Implantation



Study 1



Study 2



*All patients, including placebo-treated patients, received standard preoperative mydriatics and anesthetics

Pupil Diameter < 6 mm is Associated with a Multiply Increased Rate of Complications (e.g., Posterior Capsule Tears, Retained Lens Fragments, and Vitreous Loss)



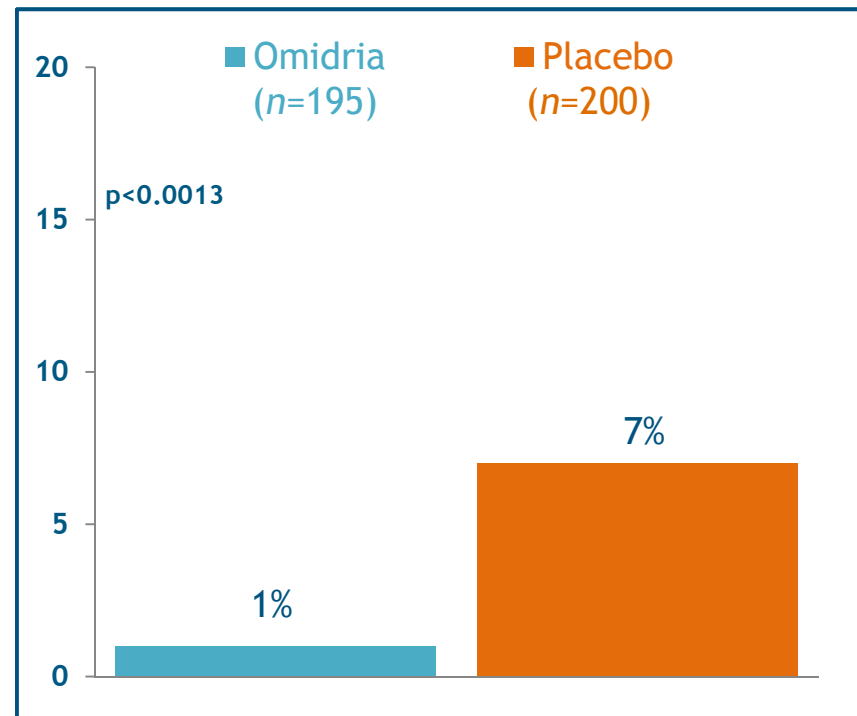
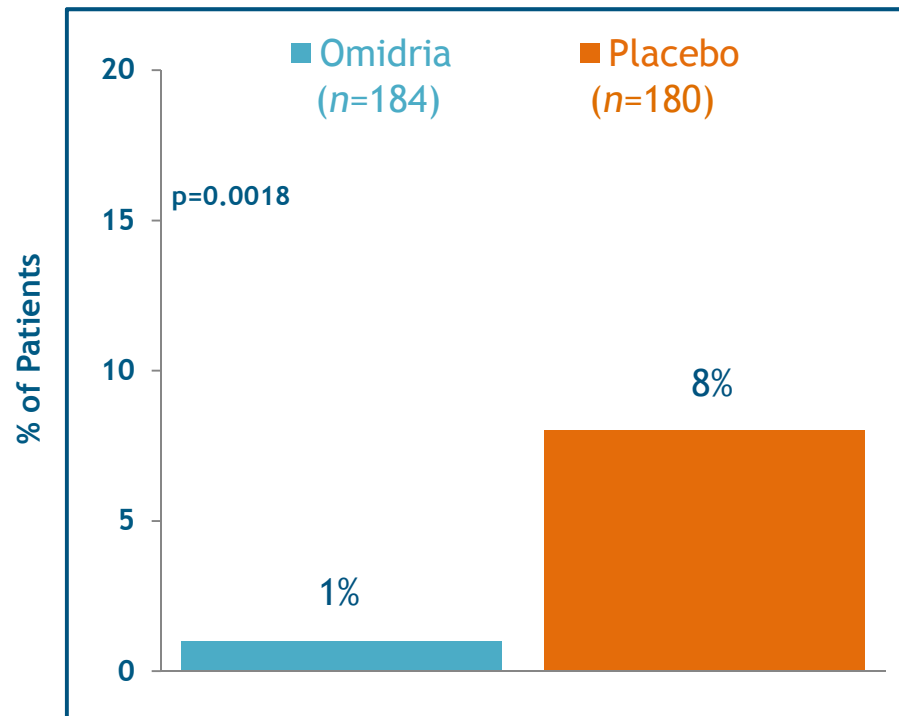
Omidria™ Phase 3 Trials

Intraoperative Miosis - Pupil Diameter < 4 mm



Study 1

Study 2



Fewer Than 1% of Omidria Subjects Experienced Pupil Diameter < 4 mm

*All subjects, including placebo-treated subjects, received standard preoperative mydriatics and anesthetics

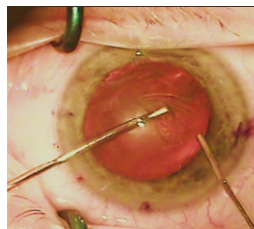
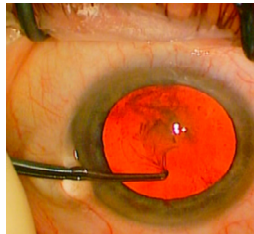


Omidria™ Importance of a Well-Dilated Pupil

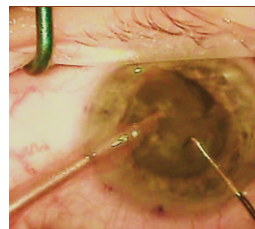
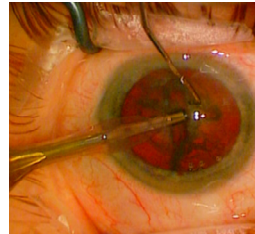


Despite Every Advance (e.g., phaco, ECCE, femto) in Cataract Surgery Over the Past Four Decades, *Small Pupils are Still a Factor Associated with Cataract Surgery Complications*

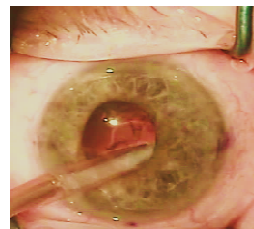
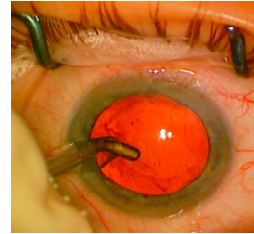
Capsulorrhexis



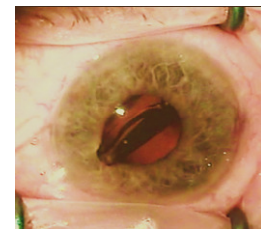
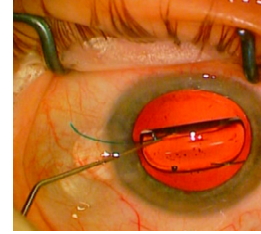
Phacoemulsification



Cortical Clean-Up



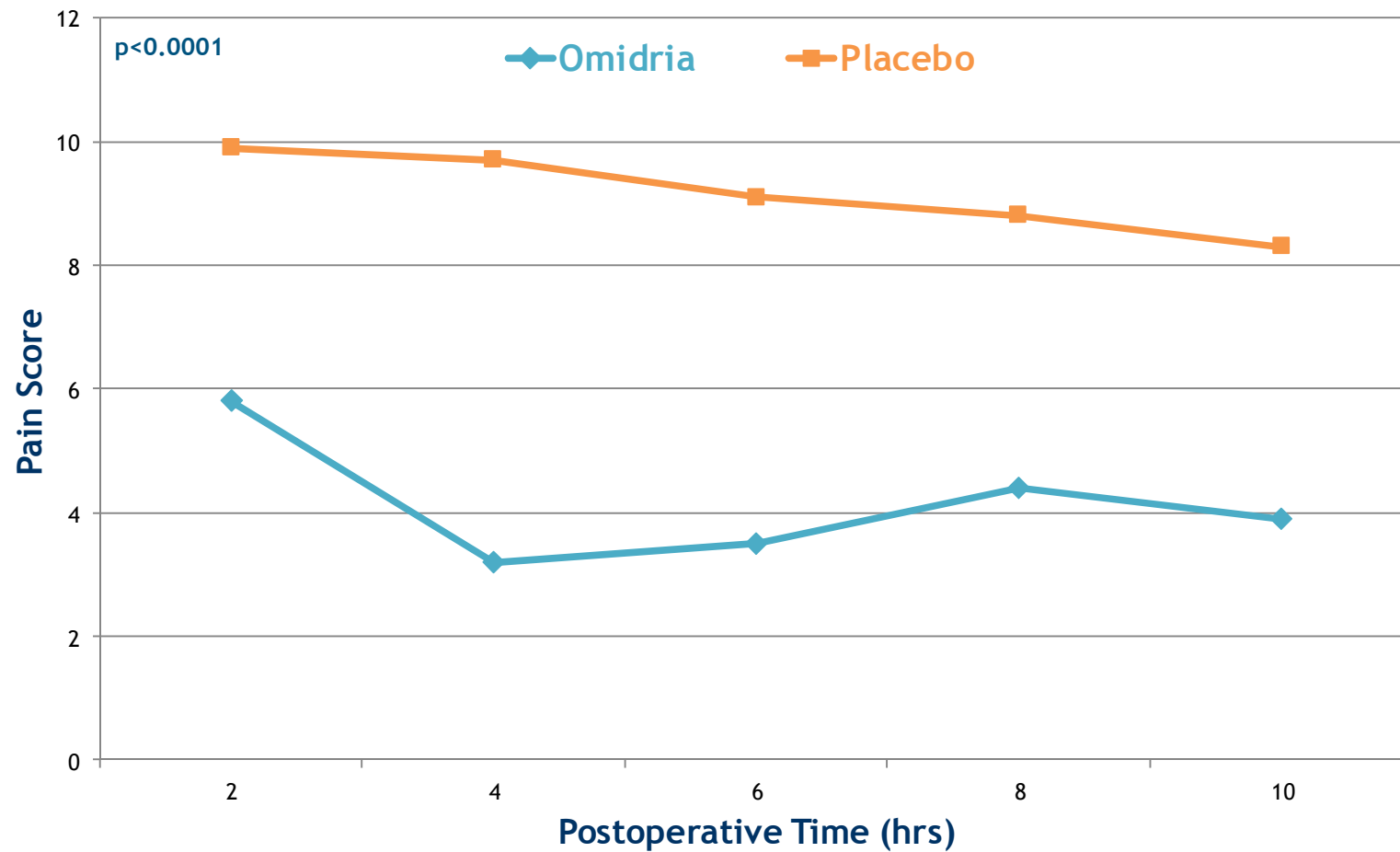
Lens Implantation



Poor Visualization Increases the Risk of Complications



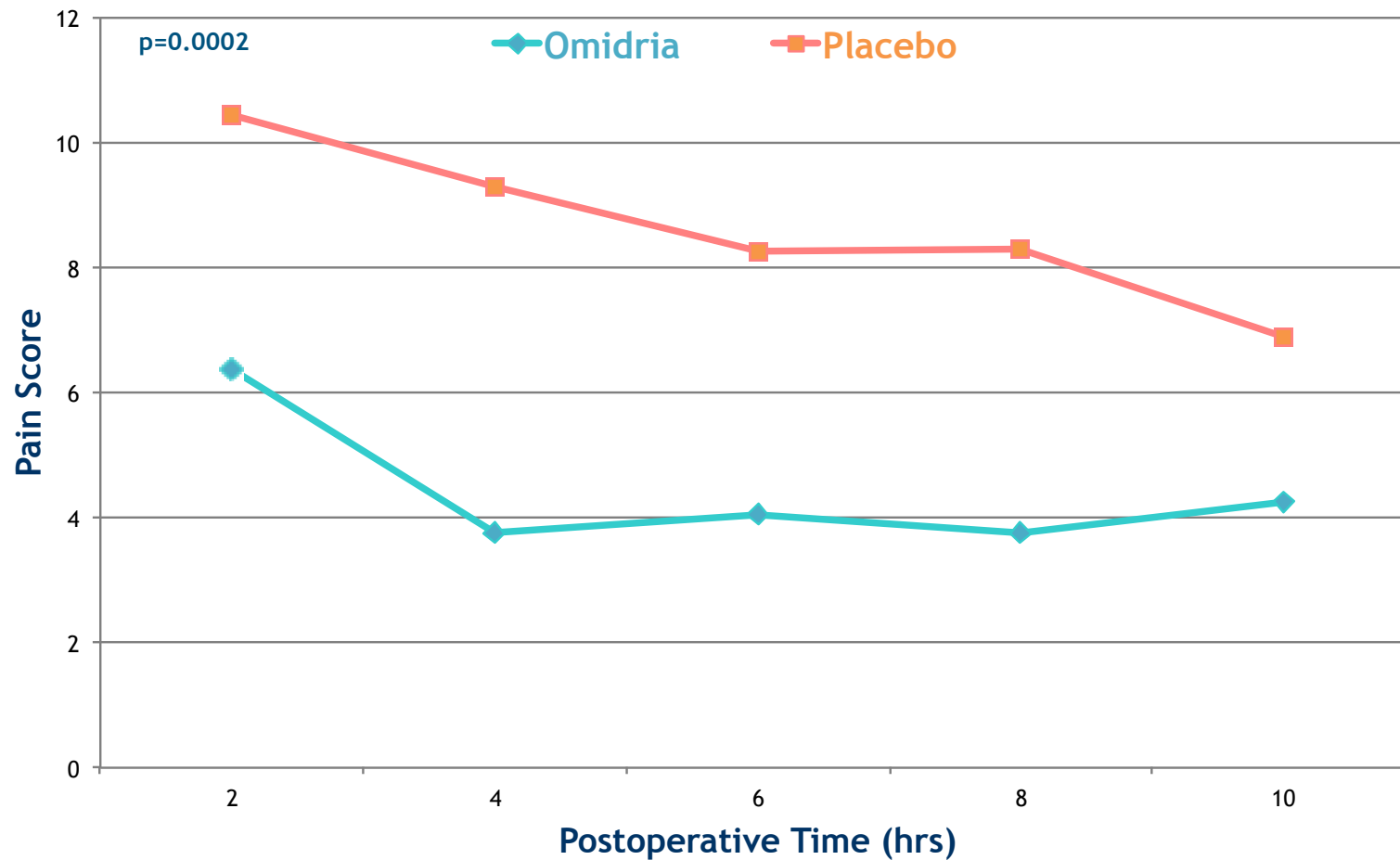
Omidria™ Phase 3 - Study 1 Postoperative Pain



*All patients received standardized preoperative mydriatics and anesthetics.



Omidria™ Phase 3 - Study 2 Postoperative Pain



*All patients received standardized preoperative mydriatics and anesthetics.

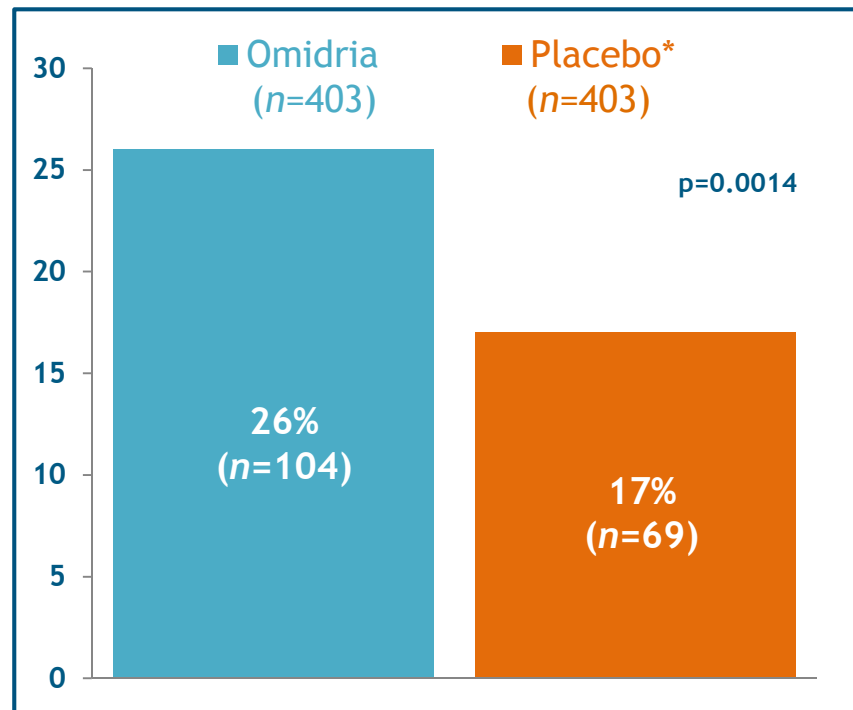
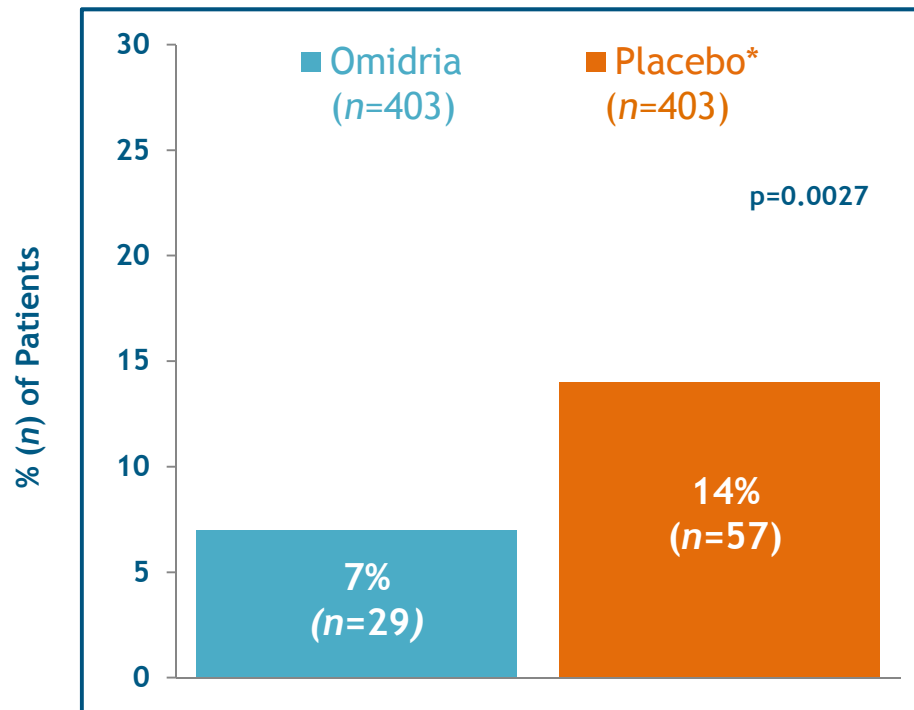


Omidria™ Phase 3 Trials Early Postoperative Ocular Pain



Moderate-to-Severe Pain (VAS ≥ 40 At Any Time Point)

Pain-Free Patients (VAS = 0 At All Time Points)



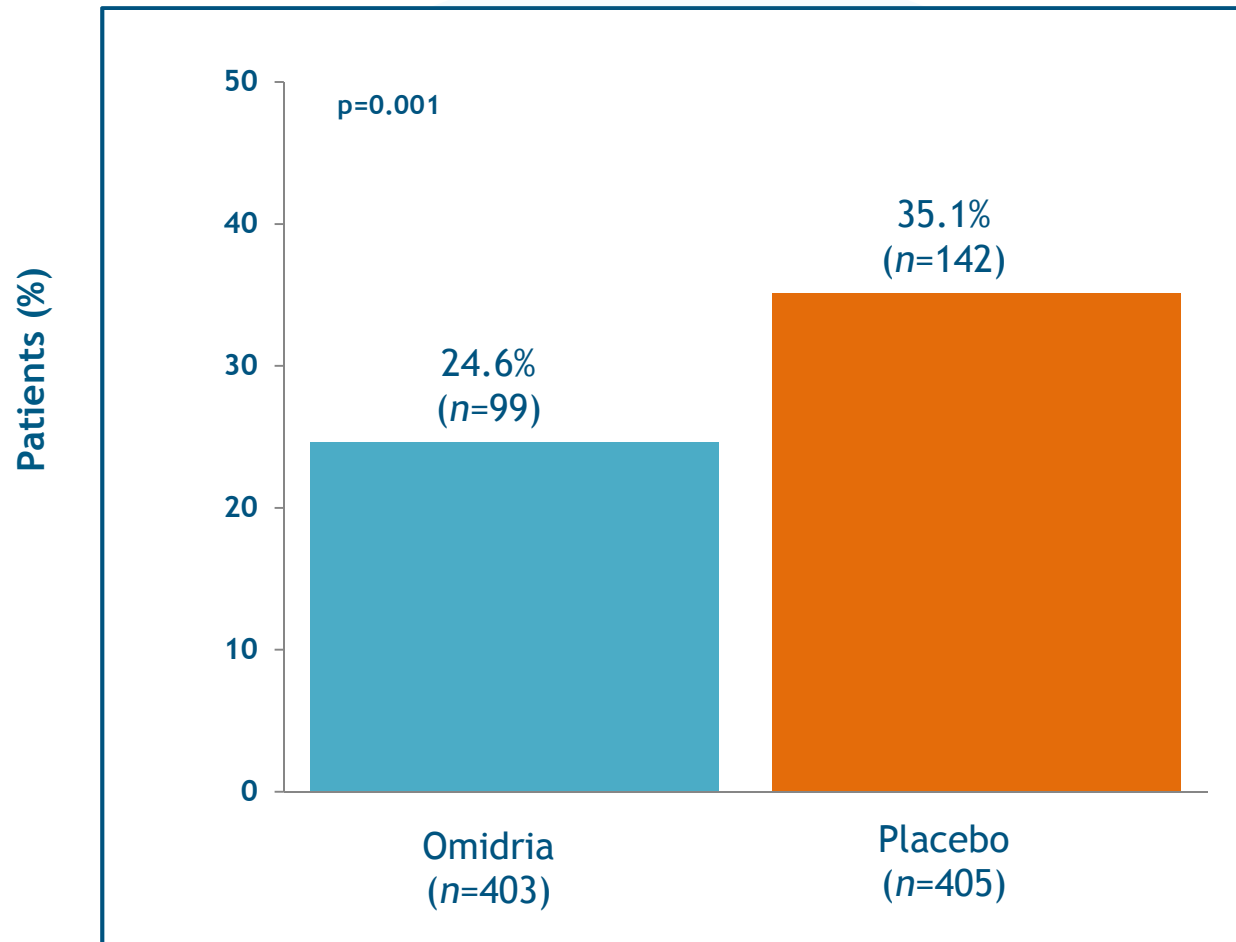
*All patients received standardized preoperative mydriatics and anesthetics.



Omidria™ Phase 3 Results Pain Medication Use



Analgesic Use on Day of Surgery



Post hoc analysis



Omidria™ Phase 3 Trials Summary



Intraocular Lens Replacement Surgery

Efficacy

- Clinically Meaningful Improvement in:
 - Maintenance of Mydriasis and Prevention of Miosis
 - Postoperative Pain Reduction
- 580% More Placebo Patients than Omidria Patients Experienced Pupil Size of < 6 mm at Cortical Clean-up
- 197% More Placebo Patients than Omidria Patients Reported Moderate-to-Severe Pain in the Early Postoperative Period
- 51% More Omidria Patients than Placebo Patients Reported No Pain in the Early Postoperative Period

Safety

- Omidria was Well Tolerated
- Adverse Events Comparable Between Omidria and Placebo
- Most Common Ocular Adverse Events were Eye Pain and Inflammation, and Increased IOP



Regulatory

- FDA Approval in May 2014
- FDA Approved Broadly for Use in All Lens Replacement Procedures (e.g., Cataract with Standard or Premium Lens; RLE)
- US and European Pediatric Plans Approved
- MAA under Review - Centralized Procedure
- European Approval Expected in 2015

Market

- 4M IOL Replacement Procedures in US, 23M Worldwide in 2015*
- Annual Growth Rate 3-4%*
- US and European KOL Advisory Boards Established
- First and Only FDA-Approved Product for Intraocular Use to Maintain Pupil Dilation and to Reduce Postoperative Pain
- Favorable Regulatory Environment with Increasingly Stringent Policies toward Compounded Products

**2011 Comprehensive Report on the Global
IOL Market (May 2011)*



Omidria™ Reimbursement & Launch



Reimbursement

- CMS Transitional Pass-through Reimbursement Granted
 - Reimbursed at WAC (\$465)+6% Followed by ASP+6%
 - Expected to Extend through December 31, 2017
- Commercial Payers May, But are Not Obligated to, Follow CMS
- Finalizing Hiring of National and Regional Reimbursement Team

US Launch

- In-House Sales, Marketing and Back-Office Management Teams
- 40 Sales Representatives Able to Access:
 - >80% of Medicare Cataract Procedures
 - >80% of Top 50, 100 and 150 Cataract Surgeons by Volume
- Commercial Manufacturing and Distribution Established
- Successful Controlled Launch in February/March
- Broad Launch Underway -- Coincides with ASCRS Annual Meeting

European Launch

- Launch Expected in Late 2015, Assuming EMA Approval Followed by Partnering



Omidria™ Barriers to Entry



Patents

- Directed to Intraoperative Delivery to the Eye of Any Combination of Agents Drawn from Two or More of the Following Classes:
 - Mydriatic
 - Anti-Inflammatory
 - Analgesic / Local Anesthetic
 - Intraocular Pressure Lowering
- 31 Issued / 15 Pending
- Method and Composition Claims Granted in US and Europe
- Excluding Any Extensions, Issued Patents Expire in 2023, Pending Applications Expire in 2033

New Legislation - DQSA

- Traditional Compounding Pharmacies (503A) Require Prescriptions
- FDA Increasingly Prohibiting Compounding at Surgery Centers
- Compounders Registered as Outsourcing Facilities (503B) Must Pay Fees, Follow GMP, Undergo FDA Inspection and Enforcement
- Neither 503A nor 503B Compounders May Compound Bulk Quantities of Close Copies of Approved Products
- Outsourcing Facilities and Traditional Compounders are Subject to Patent Enforcement with Potential Treble Damages



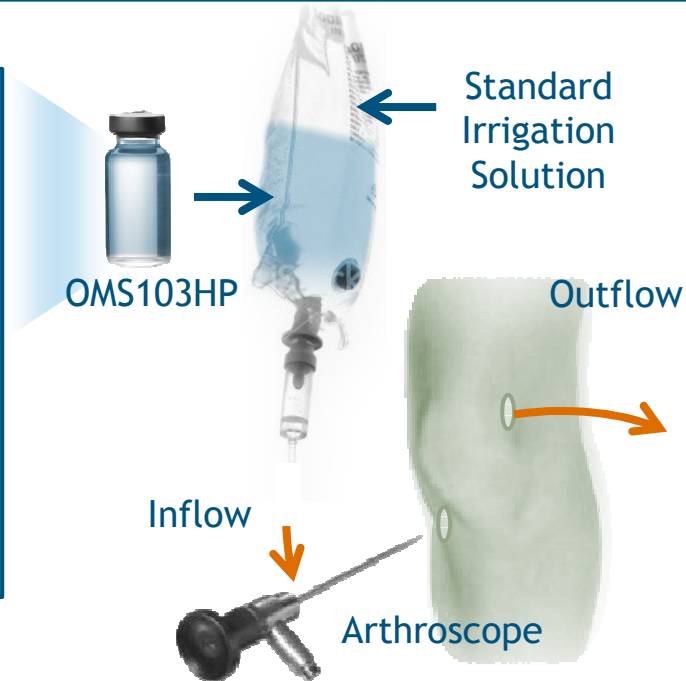
OMS103HP: Arthroscopy

Meniscectomy

4.4M Arthroscopic Procedures in US Annually

Proprietary Combination of:

- Ketoprofen
(COX-1/COX-2 Inhibitor)
- Amitriptyline
(Inhibitor of Biogenic Amines)
- Oxymetazoline
(Vasoconstrictor and Inhibitor of Neurogenic Inflammation)



Potential Clinical Benefits

- Reduce Postoperative Pain
- Improve Recovery of Range of Motion
- Improve Recovery of Joint Function
- Reduce Detrimental Inflammatory Effects on Long-Term Joint Health

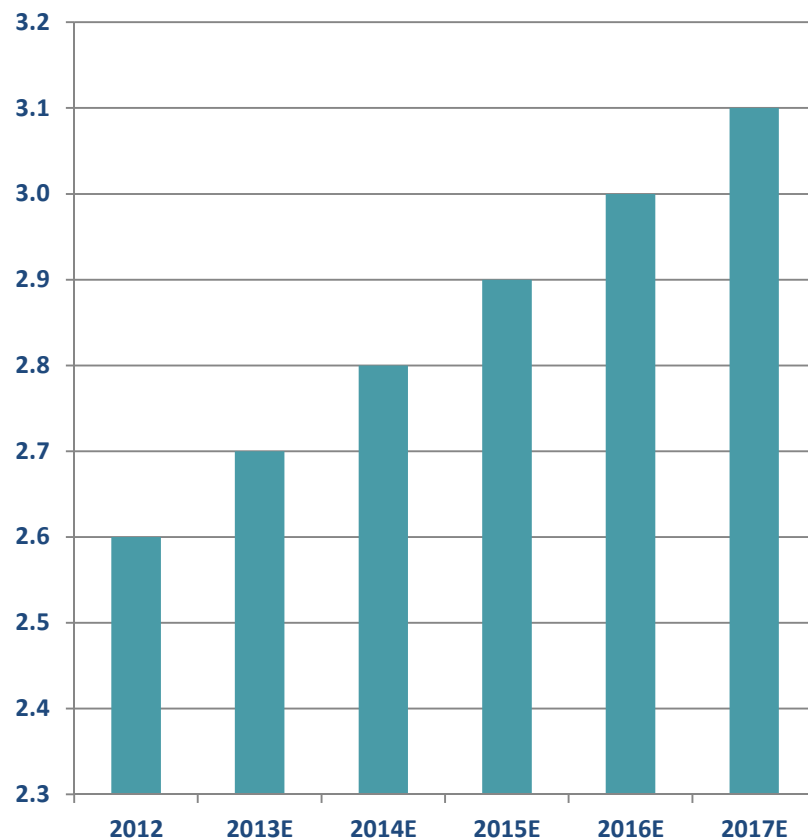


OMS103HP: Arthroscopic Market Opportunity



US Knee Arthroscopy Procedures

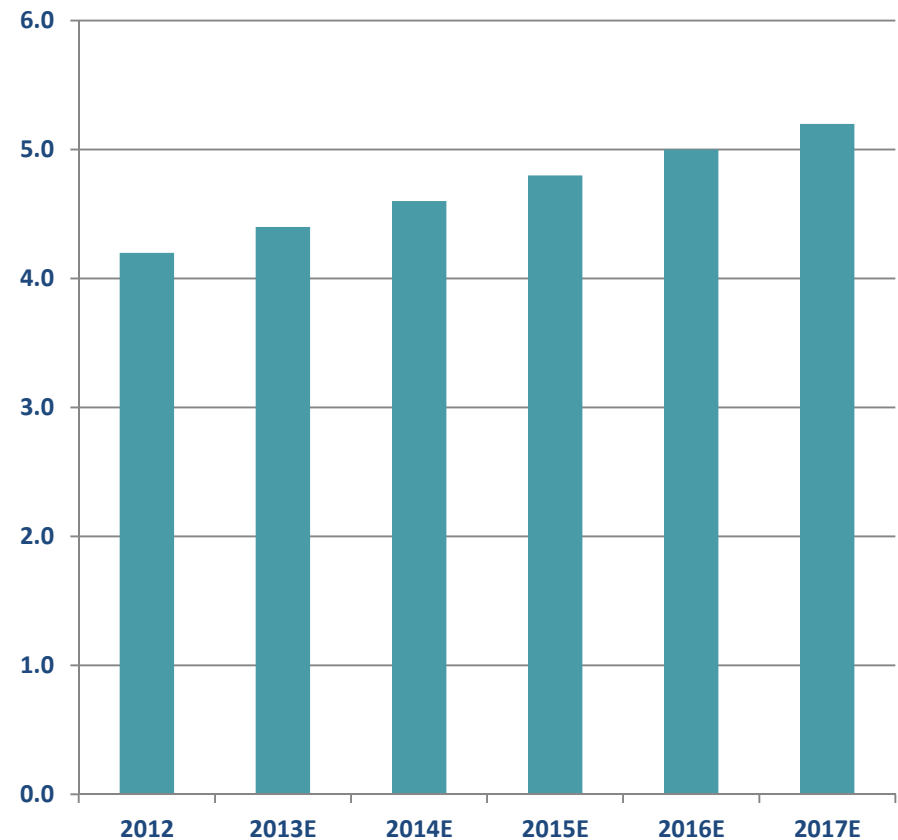
(# of Procedures in Millions)



Source: Sharon O'Reilly Consulting

US Total Arthroscopy Procedures

(# of Procedures in Millions)





OMS103HP: Meniscectomy Phase 2 Trial



Summary

Trial Design

- Vehicle-Controlled, Randomized, Double-Blind Study
- Multicenter (10 US Sites)
- Two Arms: OMS103HP and Vehicle
- 90-Day Postoperative Follow-Up
- Full Analysis Population = 143

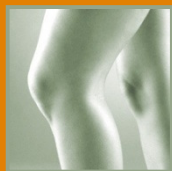
Key Endpoints

- Pain
- Knee Function
- Range of Motion

Study Results*

- Clinically Meaningful Improvement in:
 - Pain
 - Function
 - Range of Motion
- Consistent Over 90 Days
- Safe and Well-Tolerated

**The Journal of Arthroscopic and Related Surgery,
Vol. 27, No. 8, August 2011*



OMS103HP: Meniscectomy Phase 3 Trial - MEN-002



Overview

Trial Design

- Vehicle-Controlled, Randomized, Double-Blind Study
- Multicenter (18 US Sites)
- Two Arms: OMS103HP and Vehicle
- 30-Day Efficacy Postoperative Follow-Up
- Full Analysis Population = 308

Key Endpoints

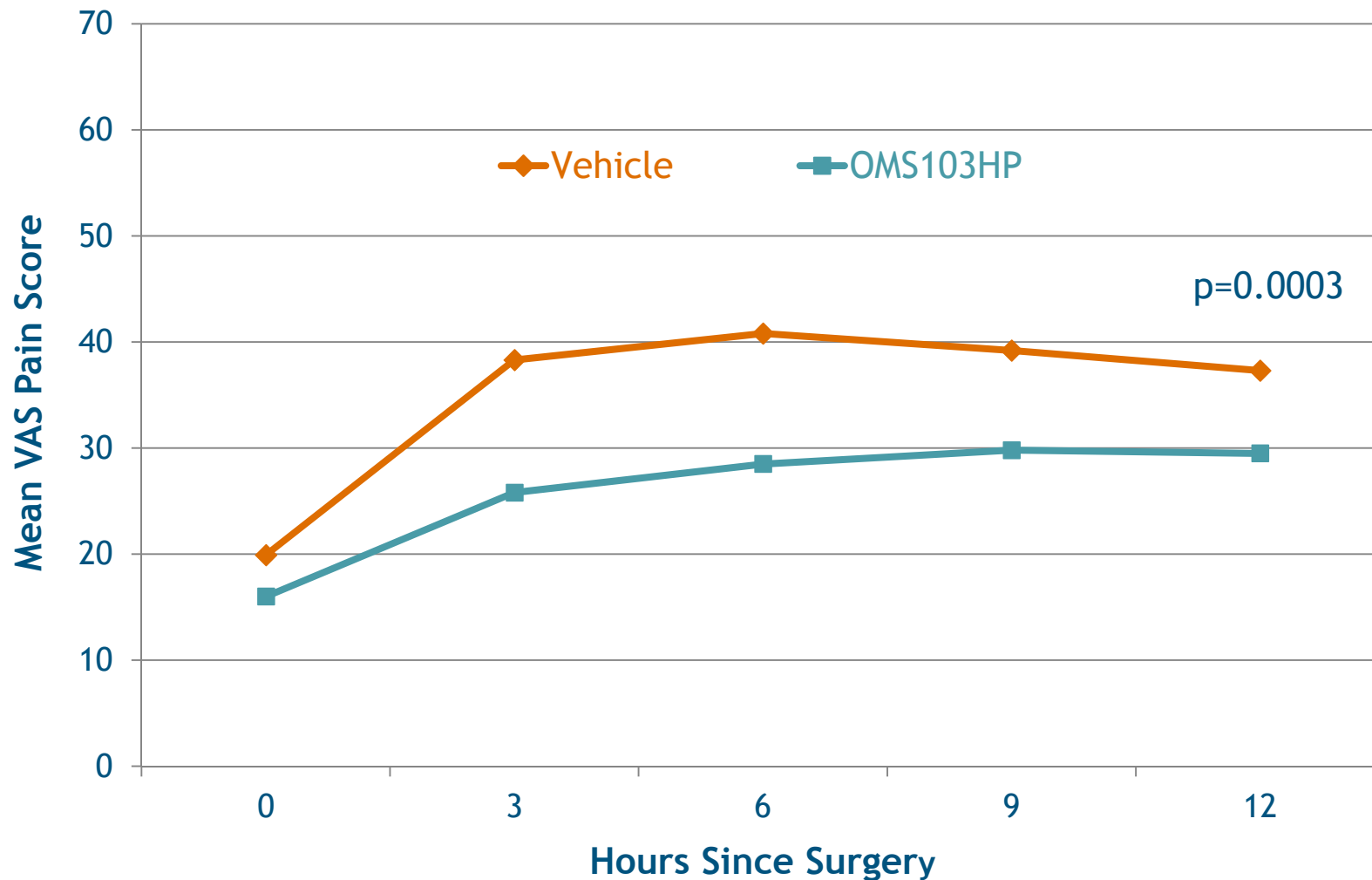
- Knee Symptoms
- Pain
- Range of Motion

Study Results

- No Significant Difference in the KOOS Symptoms Subscale
- Statistically Significant and Clinically Meaningful
 - Improvement in Early Postoperative Pain
 - Decreased Postoperative Opioid Use
- Incidence of Inflammatory Adverse Events, Tourniquet Use and Crutch Use, as Well as Time to Discontinuation of Crutches and Return to Work All Favored OMS103HP
- Safe and Well-Tolerated

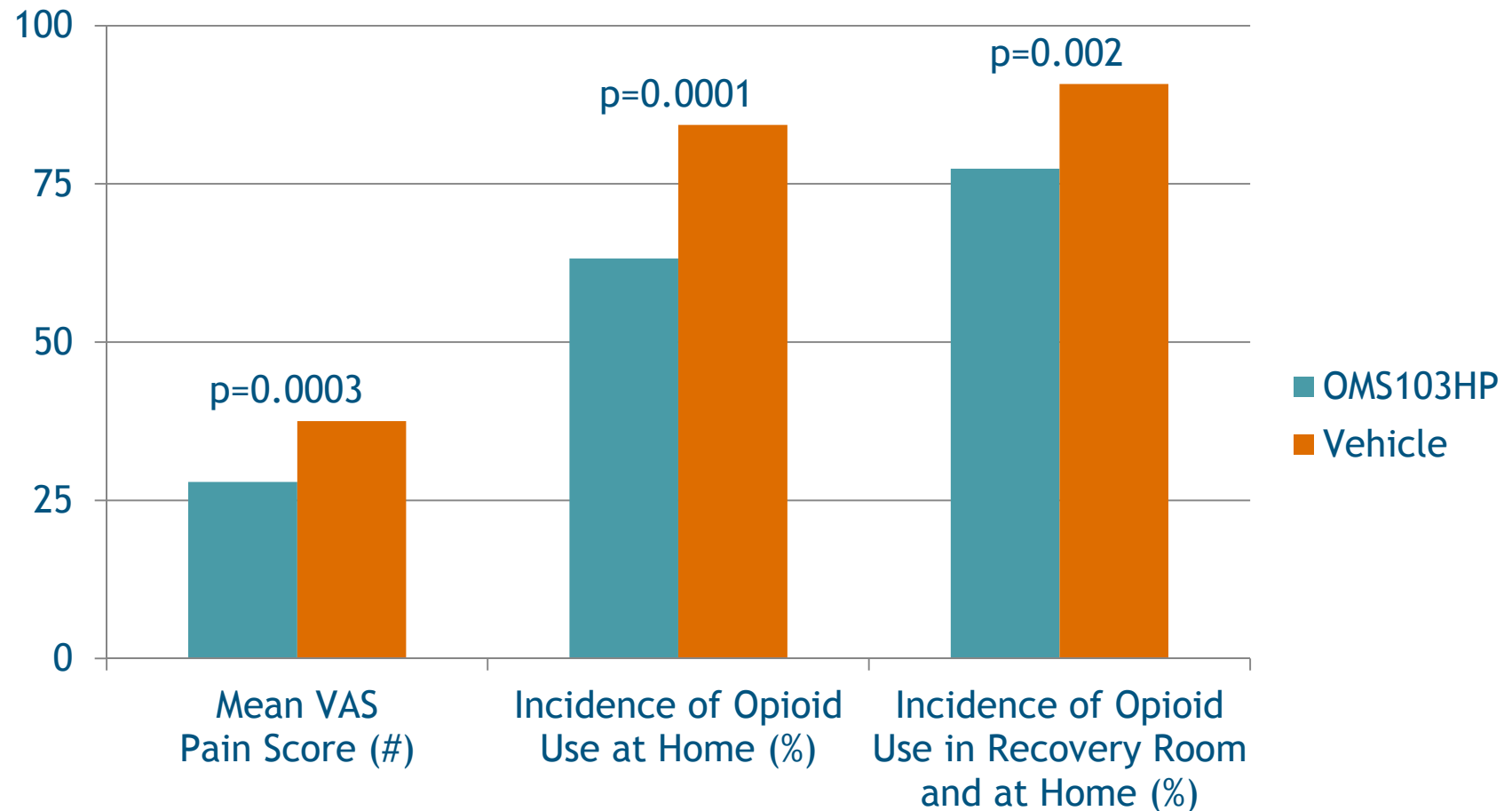


OMS103HP: Phase 3 Meniscectomy MEN-002 - Early Postoperative Pain Scores





OMS103HP: Phase 3 Meniscectomy MEN-002 - Postoperative Pain and Opioid Use





OMS103HP: Phase 3 Meniscectomy Study Conclusions



Efficacy

- Clinically Meaningful Improvement in Early Postoperative Pain
- 134% More OMS103HP Patients Required No Postoperative Opioids at Home on the Day of Surgery
- 146% More OMS103HP Patients Required No Postoperative Opioids on the Day of Surgery
- While Not Powered for Significance, More OMS103HP Patients:
 - Reported No Postoperative Inflammatory Adverse Event
 - Did Not Use Crutches
 - Reported Early Discontinuation of Crutch Use
 - Did Not Have a Tourniquet Used During the Procedure
- No Significant Difference in KOOS Subscales

Safety

- Incidence of Adverse Events Was Higher with Vehicle
- Safe and Well-Tolerated

Next Steps

- Re-designing Phase 3 Program to Make Pain the Primary Endpoint

Alternative Commercialization Opportunity for PharmacoSurgery® Platform



Manufacture and Sales through Outsourcing Facilities

Compounding Quality Act

- Title I of Drug Quality and Security Act Passed Nov 2013
- Reaffirmed Rules for Traditional Compounding Pharmacies (Section 503A) - Individual Patient Prescriptions Required
- Created “Outsourcing Facilities” (Section 503B)

Outsourcing Facilities

- Must Register with FDA and Follow Good Manufacturing Practices (GMP) for All Compounded Products
- May Produce “in Bulk” without Need for Prescriptions
- May Advertise Using Clinical Data if Not False or Misleading

Revenue Opportunity

- PharmacoSurgery® Products (e.g., OMS103 and OMS201) are Broadly Patented Combinations of Existing APIs
- Formulation, Manufacturing and Stability are Well Established
- Compelling Clinical Data from Placebo-Controlled Trials
- No Additional Clinical Trials Required



Pipeline Programs

OMS721: MASP-2 Antibody



Overview

- Specifically Blocks the Lectin Pathway (LP)
- Omeros Controls Worldwide Exclusive Rights to the Inhibition of MASP-2 and to All MASP-2 Inhibitors
- FDA Orphan Designation: Prevention of Complement-Mediated TMAs

OMS721 vs. Soliris®

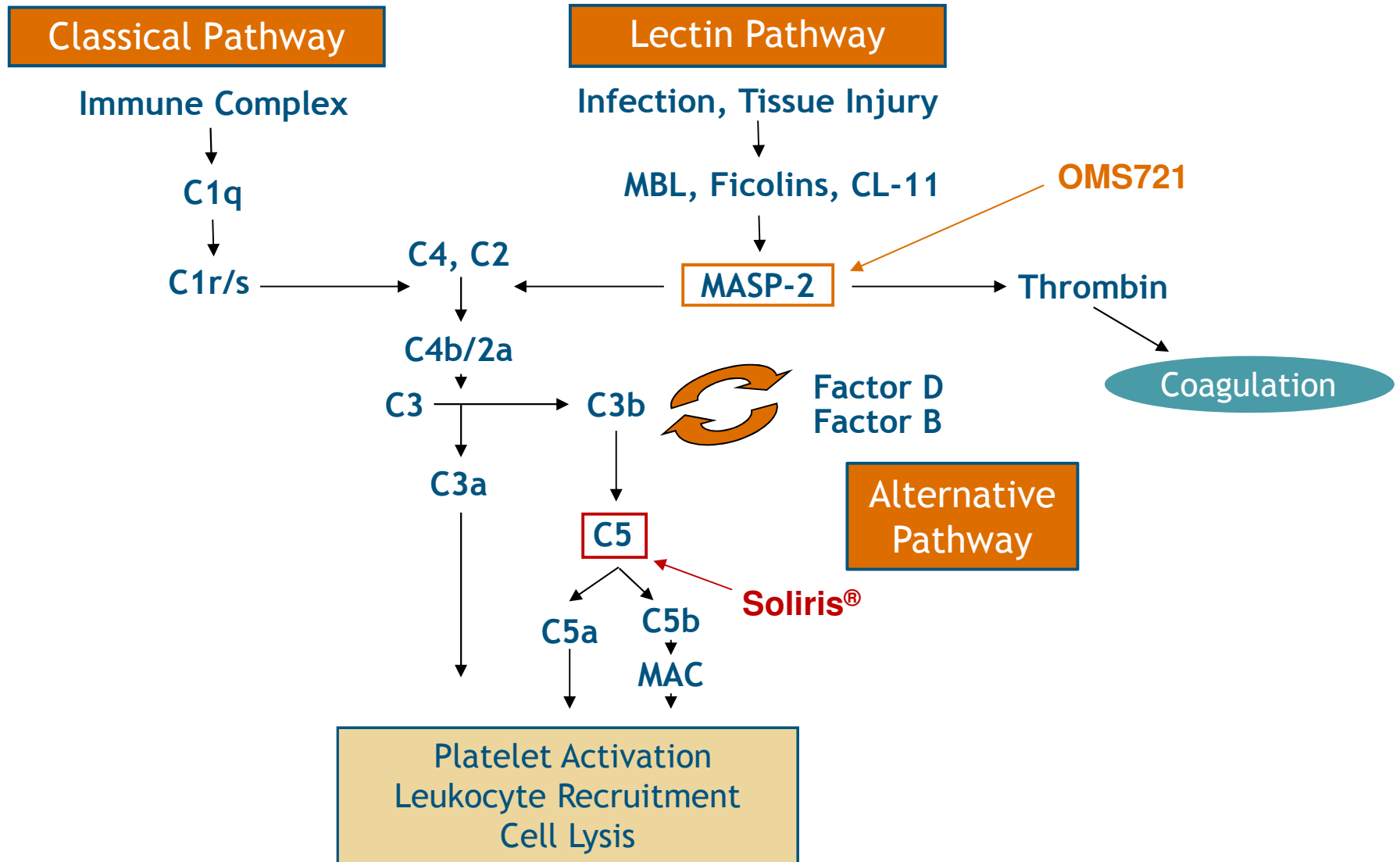
- Classical Pathway Unaffected
- Subcutaneous Dosing vs. Intravenous Infusion
- Role in Coagulation Cascade

Expanding Patent Position

- Identified MASP-3 as Key Activator of Alternative Pathway
- Advancing High-Affinity, Functionally Active MASP-3 mAbs
- Generated *In Vivo* Data around Multiple Indications
- Expands IP Position to Both Lectin and Alternative Pathways



OMS721 Targets MASP-2: Protease Required for Lectin Pathway Function





OMS721 Clinical Trials



Phase 1 Clinical Program

- Evaluated Safety, Tolerability, Pharmacodynamics and Pharmacokinetics of OMS721 in Healthy Subjects
- Well Tolerated by Both Intravenous and Subcutaneous Administration
- Subcutaneous Administration Yielded Sustained LP Inhibition

Phase 2 Clinical Program

- Phase 2 Clinical Trial Evaluating OMS721 in Patients with Complement-Mediated Thrombotic Microangiopathies
 - Includes aHUS, TTP, Stem Cell Transplant-Related TMA
- Improvements Seen Across all Disease Markers In Low-Dose Cohort
- Compassionate Use Granted for Two Patients in Low-Dose Cohort
- Dosing Complete for Mid-Dose Cohort
- Dose Escalation to High-Dose Cohort Planned for this Month
- Now Able to Dose Chronically in Humans
 - No Drug-Related Adverse Events in Chronic Toxicology Study
- Additional Phase 2 Data Expected Later this Year



OMS721

Ex Vivo Evaluation in aHUS Patient Serum

Background

- Exposure of Human Microvascular Endothelial Cells (HMEC-1) to aHUS Patient Serum, But Not Control Serum, Leads to Complement Activation and Formation of Thrombus
- Complement Deposition and Thrombus Formation are Blocked By Co-incubation of Patient Sera with Eculizumab
- Complement Deposition Response Normalizes in aHUS Patients on Eculizumab Therapy Concurrent with Clinical Remission*

Objectives

- Test effect of OMS721 on aHUS Patient Serum-induced:
 - Complement Deposition
 - Thrombus Formation

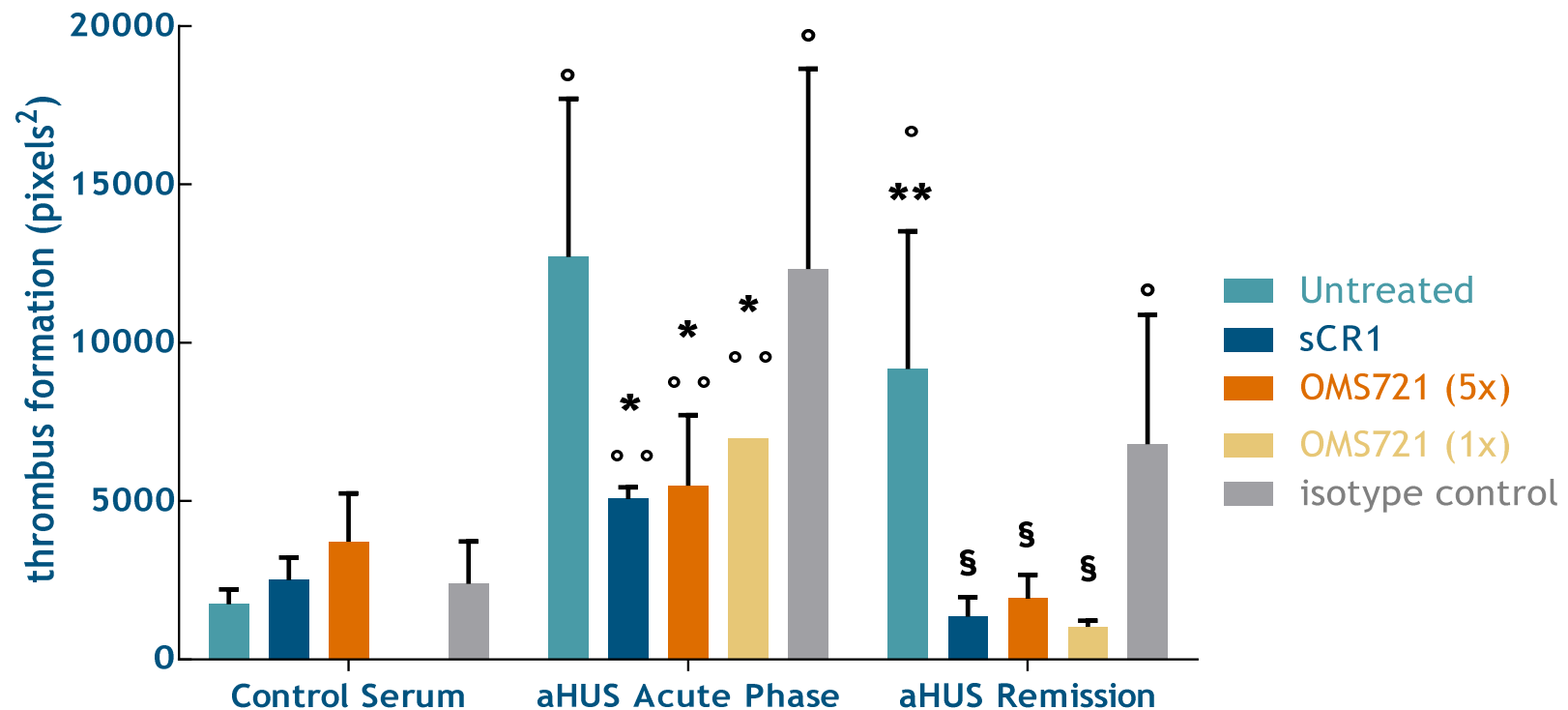
Results

- Complement Deposition: OMS721 Effect Equal to Published Effect of C5 Inhibitors
- Thrombus Formation: OMS721 Effect Equal to that of Positive Control (SCR-1)

*Gastoldi et al., Immunobiology (2012) 217:1145
Noris et al., Blood (2014) 124:1715



OMS721 Blocks aHUS Serum-Stimulated Thrombus Formation



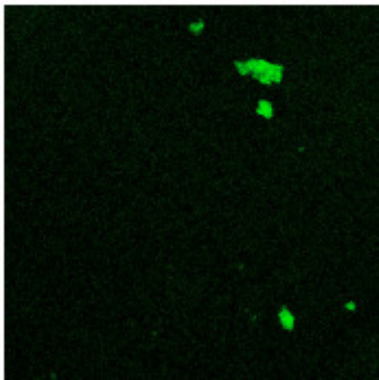
aHUS serum-induced thrombus formation on human microvascular endothelial cells. Data are mean \pm SE.
[°] P<0.0001, ^{°°} P<0.01 vs control; *P<0.0001, **P<0.01 vs aHUS acute phase untreated; § P<0.0001 vs aHUS remission phase untreated.

Unpublished data: Remuzzi, et al. 2014

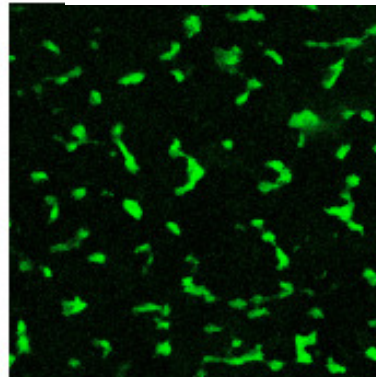


OMS721: Representative Images of aHUS Serum-Stimulated Thrombus Formation

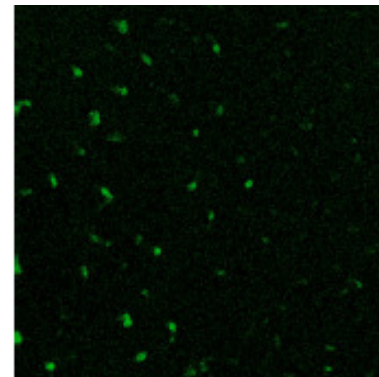
Normal Healthy



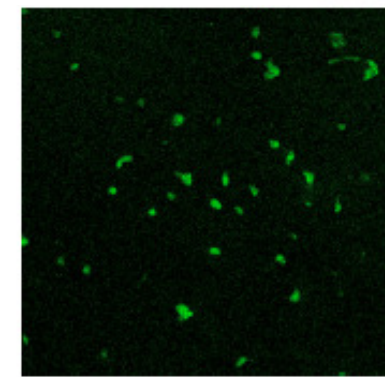
Untreated



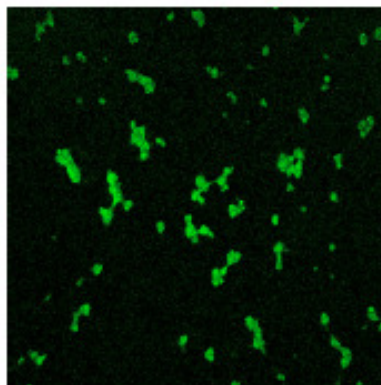
aHUS Acute Phase
+ sCR1



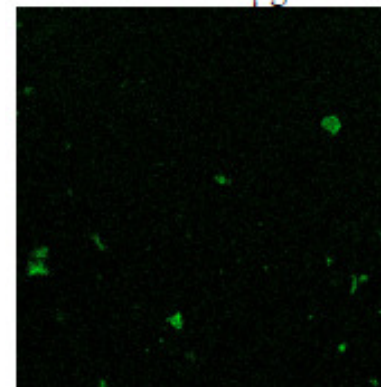
+ OMS721



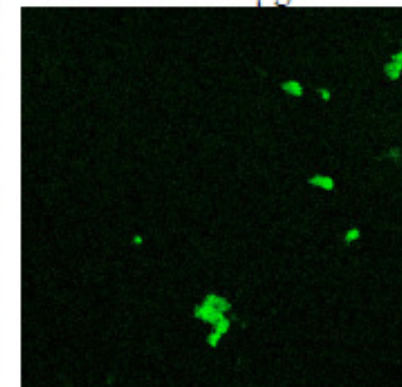
Untreated



aHUS Remission
+ sCR1



+ OMS721

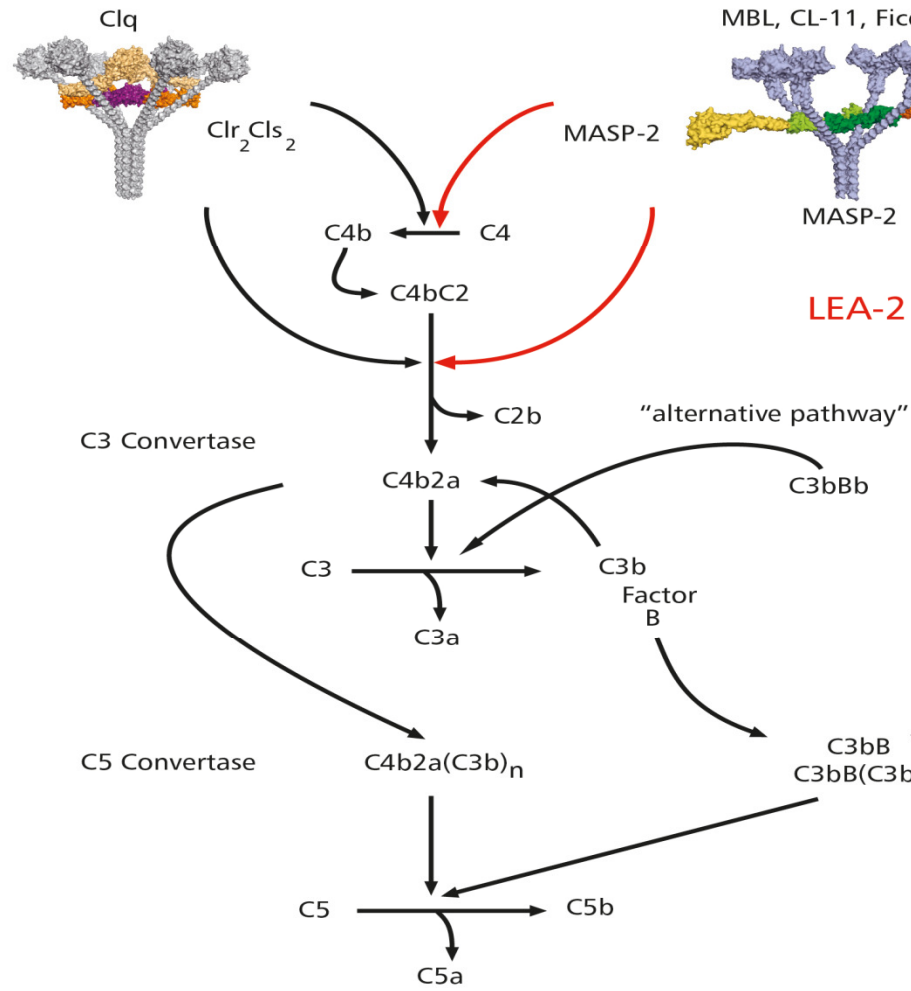




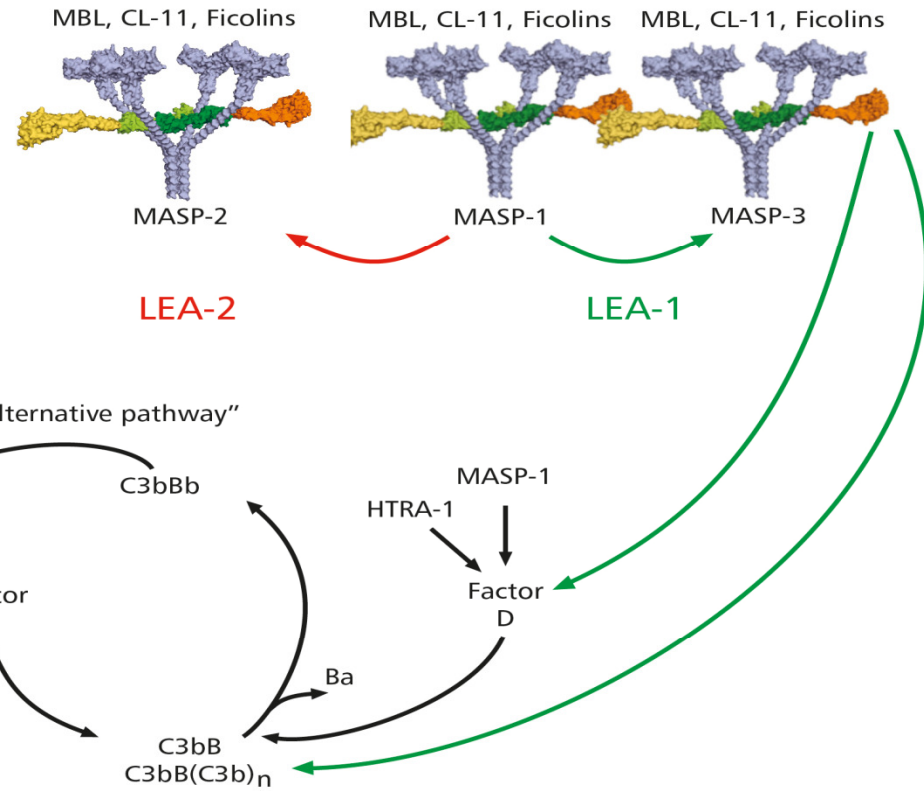
OMS721

Lectin Pathway Effector Arms (LEA)-1 and -2

Classical Pathway











Lectin Pathway





OMS721 is the Only Lectin Pathway-Specific Complement Inhibitor in DevelopmentX



Product	Target	Pathways Blocked	Indication(s)*	Limitations	Status
 OMEROS OMS721	MASP-2	Lectin	TMA (Including aHUS), AMD, DN, MN, MI, Stroke, Transplant		Phase 2
 ALEXION Soliris®	C5	Alternative, Classical & Lectin	aHUS, PNH, NMO, MG, CAD, Transplant	1. High Dosing Required 2. IV Infusion Required 3. Safety Concerns	Marketed
 NOVARTIS POT-4	C3	Alternative, Classical & Lectin	AMD	1. No Systemic Delivery Option	Phase 2
 ALEXION ALXN1103	C3b	Alternative	PNH	1. High Dosing Required 2. Short Half-life 3. Safety Concerns	Phase 1
 NOVARTIS LFG316	C5	Alternative, Classical & Lectin	AMD, Choroiditis	1. Intravitreal	Phase 2
 OPHTHOTECH Zimura	C5	Alternative, Classical & Lectin	AMD	1. Intravitreal	Phase 2
 Genentech <small>A Member of the Roche Group</small> Lampalizumab	Factor D	Alternative	AMD	1. Intravitreal	Phase 3
 Apellis APL-2	C3	Alternative, Classical & Lectin	PNH	1. Short Half-life 2. Safety concerns	Phase 1

*aHUS: atypical hemolytic uremic syndrome; AMD: age-related macular degeneration; CAD: cold agglutinin disease; DN: diabetic neuropathy; MG: myasthenia gravis; MI: myocardial infarction; MN: membranous nephropathy; NMO: neuromyelitis optica; PNH: paroxysmal nocturnal hemoglobinuria; TMA: thrombotic microangiopathy



Pipeline Programs

OMS824: PDE10 Inhibitor



Overview

- Potential for Cognitive Enhancement and Neuroprotection
 - Huntington's Disease (HD)
 - FDA Orphan Drug & Fast Track Designations Granted
 - Schizophrenia
 - Alzheimer's Disease/Mild Cognitive Impairment
- ~70% Human PDE10 Occupancy without EPS

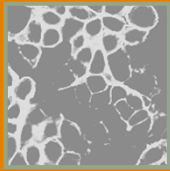
Current Indications

- HD
- Schizophrenia

Clinical Trials

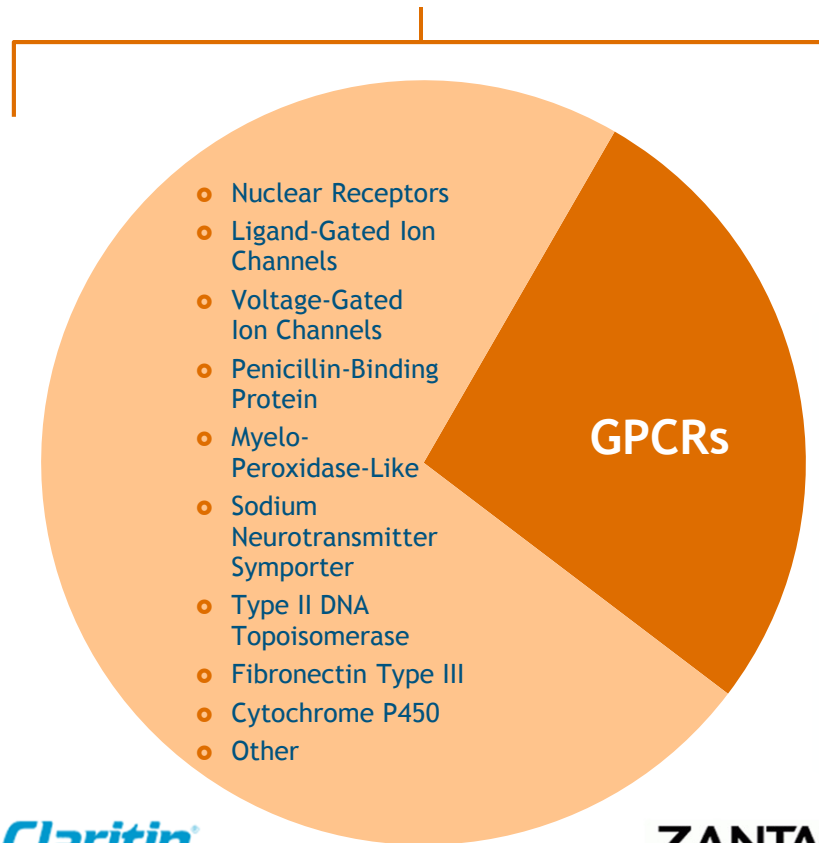
- Successful Phase 1 Single and Multiple-Ascending Dose Trials
 - Once-Daily Dosing with Linear Pharmacokinetics
 - MTD Not Yet Identified (Tolerated at Plasma Concentrations Twice that Obtained at 70% Occupancy)
- Positive Phase 2a Schizophrenia Trial
 - Dose Adjustment Not Needed When Administered with Concomitant Antipsychotics
- Phase 2 Programs Currently on Hold; Expect to Re-Initiate Enrollment in HD Program Soon

~30,000 HD Patients in US and One Approved Drug – Tetrabenazine (Chorea Only, Black Box Warning);
2.2 M Schizophrenia Patients in US / Sales of Atypical Antipsychotics >\$16B



The Significant Potential of GPCRs as Drug Targets

Annual Worldwide Drug Market
\$700B+



GPCRs: The Premier Drug Targets

Characteristics

Modulation of Numerous Physiological Procedures

High Specificity and Limited to Specific Tissues

Expressed on Cell Surface

Advantages

Impacts **Many** Diseases

Limited Side Effects

Easily Accessible by Drugs

Claritin



L-DOPA

OXYCONTIN

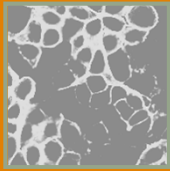
ZANTAC

ZYPREXA
Olanzapine

Lopressor

reglan

VICODIN

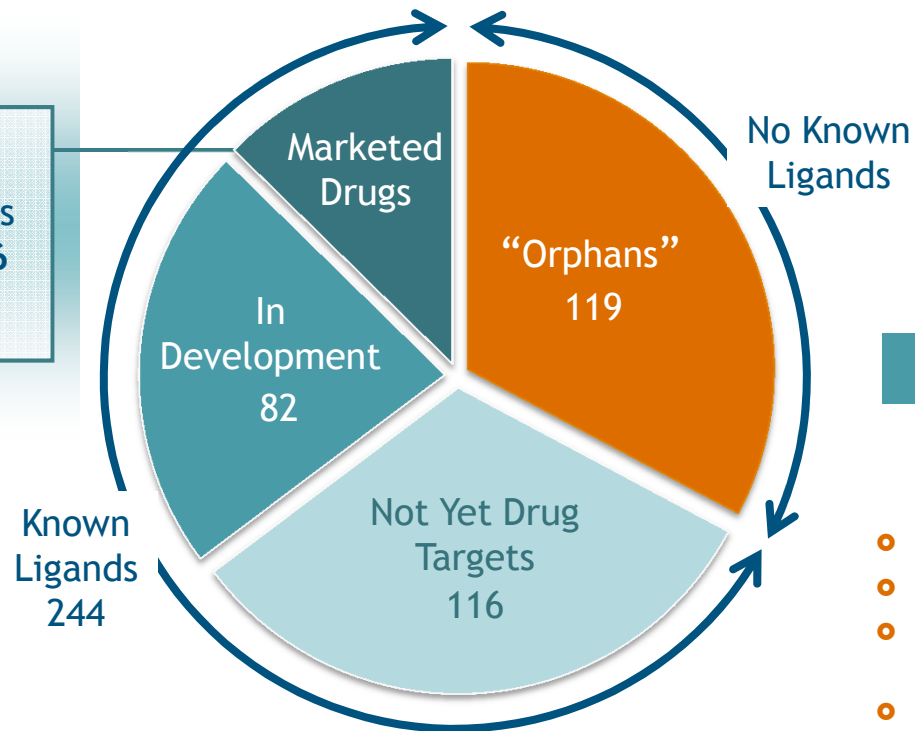


GPCRs as Drug Targets

363 Non-Sensory GPCRs

Today

30-40% of All
Marketed Drugs
Target Only 46
GPCRs



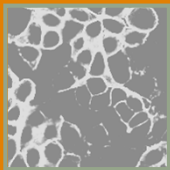
The Challenge

Promising Targets, but Drug Discovery Difficult

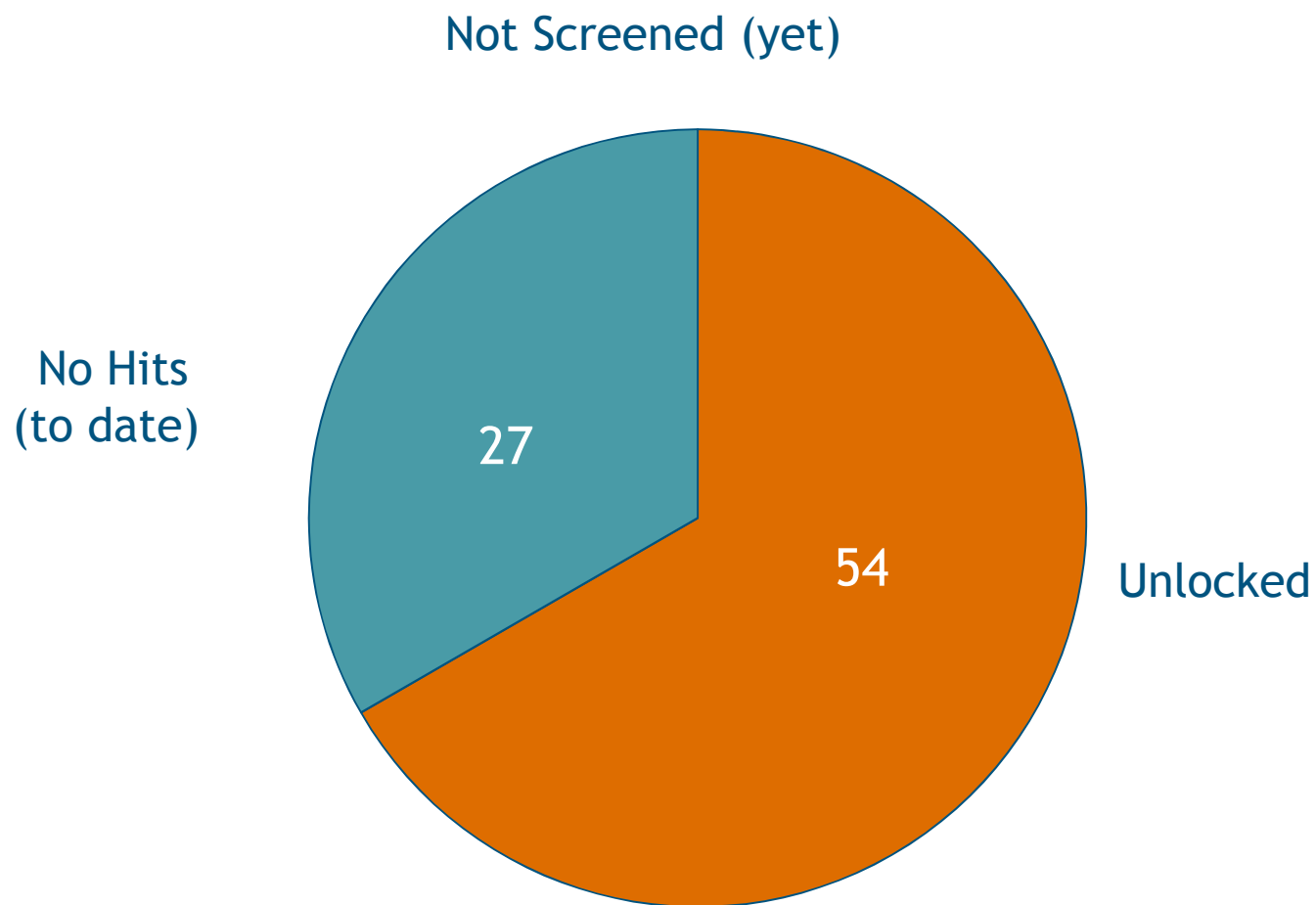
- Ligand Required for Assay Development
- Signaling Pathway Not Known
- Laborious Fractionation for Natural Ligand Identification
- Current Technologies Limited Only to Agonist Screening

The Opportunity

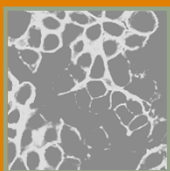
Up to 60+ New Drug Targets



Unlocking Class A oGPCRs



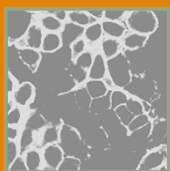
Class A oGPCRs (81)



Orphan GPCRs Unlocked To Date

GPCR	Metabolic & Cardiovascular Indications
GPR12	Obesity, Cognitive Impairments
GPR21	Obesity, Diabetes
GPR22	Cardiovascular Diseases, Anxiety
GPR25	Arterial Stiffness
GPR37L1	Hypertension
GPR39	Diabetes
GPR50	Metabolic Disorders
GPR61	Eating Disorders
GPR82	Appetite, Body Weight
GPR101	Eating Disorders
GPR132	Atherosclerosis
GPR146	Dyslipidemia, Diabetes
GPR171	Eating Disorders
GPR176	Atherosclerosis
SREB1/GPR27	Diabetes, Schizophrenia

GPCR	Oncology Indications
GPR19	Melanoma, Lung Cancer
GPR20	Gastro-Intestinal Stromal Tumors, Acute Myeloid Leukemia
GPR65	Renal Cell Carcinoma, Ovarian Cancer, Inflammation
GPR68	Ovarian Cancer, Prostate Cancer, Osteoporosis
GPR80	Hepatocellular Carcinoma
GPR87	Squamous Cell Carcinomas
GPR150	Ovarian Cancer
GPR161	Breast Cancer, Congenital Cataracts & Birth Defects
GPR174	Melanoma, Grave's Disease
LGR4	Cancer Stem Cells, Bone Diseases
LGR5	Cancer Stem Cells, Esophageal Adenocarcinoma
P2Y8	Leukemias, Lymphomas

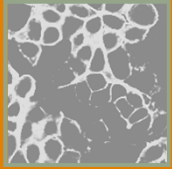


Orphan GPCRs Unlocked To Date

GPCR	CNS Indications
GPR17	Myelin Disorders, Multiple Sclerosis
GPR31	Anxiety Disorders
GPR37	Parkinson's Disease
GPR52	Schizophrenia
GPR63	Autism
GPR78	Bipolar Disorder, Schizophrenia
GPR139	Motor Disorders
GPR151	Cognition, Mood Disorders, Pain
GPR153	Schizophrenia
MAS1	Cognitive Impairments
MRGE	Pain
OPN4	Circadian Rhythm, Sleep Disorders
SREB2/GPR85	Schizophrenia, Obesity
SREB3/GPR173	Schizophrenia, Obesity

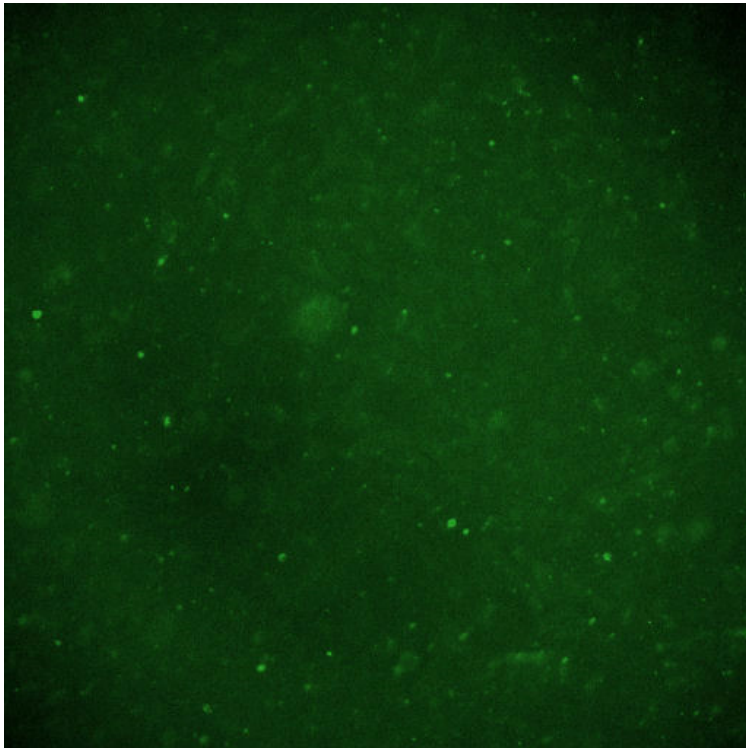
GPCR	Miscellaneous Indications
GPR15	HIV Enteropathy, Rheumatoid Arthritis
GPR32	Acute Inflammatory Responses
GPR83	Autoimmune Diseases, PTSD
GPR183	Humoral Immunity
CCRL2	Rheumatoid Arthritis
LGR6	Hair Follicle Stem Cells, Wound Repair

GPCRs with Unknown Indications	
GPR45	GPR182
GPR135	MRGF
GPR141	OPN5
GPR162	

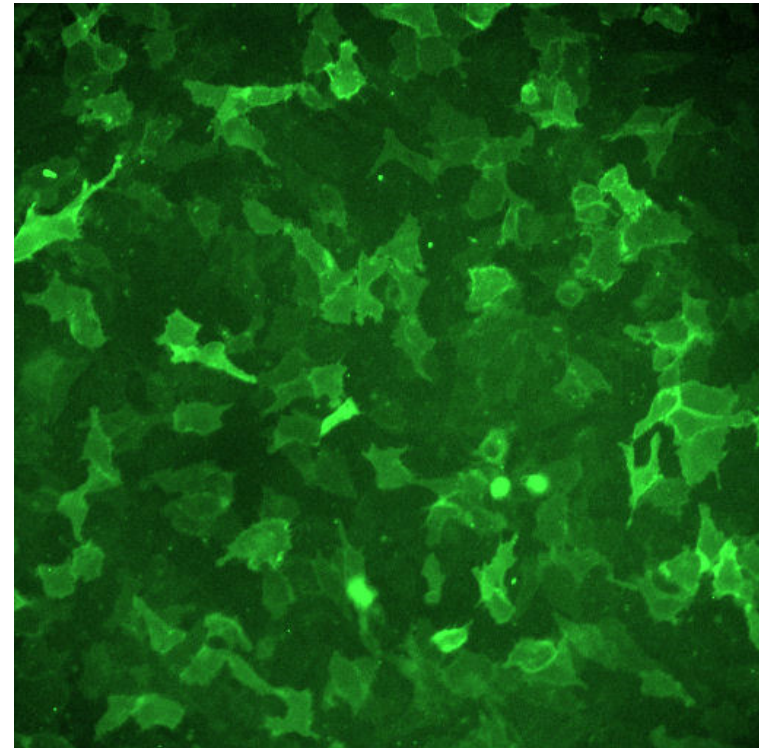


High-Throughput Cellular Redistribution Assay

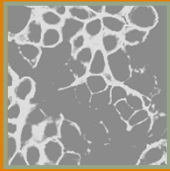
No Ligand



Ligand



Identifies Agonists, Antagonists, and Inverse Agonists



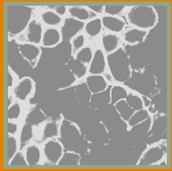
Evaluation of CRA Hits in Signaling Assays

Signaling Assays

- Reporter Assays
 - Gs, Gi, Gq, G12/13 Signaling Pathways
- GloSensor Assay
 - Gs and Gi Signaling Pathways
- Beta-Arrestin Recruitment Assay

Functional Data Provided

- Functional Confirmation of CRA Hits
- Determination of Nature of Compounds (Agonist, Antagonist, Inverse Agonist)
- Key IP Component - Uniquely Identifying Primary Signaling Profile



Establishing Intellectual Property for Orphan GPCRs

Three Levels of Intellectual Property

1. Linked to Signaling Profile

Independent of Indication

2. Linked to Knock-Out Mice/Phenotypes

Indication-Specific

3. Medicinal Chemistry*

Composition of Matter

*Also applies to establishment of IP position for recalcitrant non-orphan GPCRs



Pipeline Programs PDE7 & Plasmin



PDE7 (OMS527)




- IND/CTA Submission Planned for Late 2015/1H 2016
- Initial Indication: Cocaine/Nicotine Addiction
- No Current Drug for Cocaine Addiction
- Omeros Discovered Link between PDE7 and Addiction and between PDE7 and Movement Disorders
- Omeros Controls PDE7 Broadly for
 - Addiction Disorders and Compulsive Behaviors
 - Movement Disorders (e.g., Parkinson's Disease)

Plasmin (OMS616)

- IND/CTA Submission Planned for 2016
- Initial Indications: Trauma and Cardiovascular Surgery
- Efficacy Equal to Trasyolol® by Human *Ex Vivo* Data
- Safety Profile Potentially Superior to Trasyolol
 - Human Protein (Trasyolol is Bovine)
 - Unlike Trasyolol, No Activity at Kallikrein or Factor Xla

Products and Programs



Program	Targeted Procedure/ Disease	Pre-clinical	Phase 1	Phase 2	Phase 3	FDA Approval	Commercial Launch	Economic Rights
Commercial Products								
Omidria™ (OMS302) - Ophthalmology	Cataract Surgery or Intraocular Lens Replacement	<div></div>						<div> OMEROS</div>
Clinical Programs								
OMS103 - Arthroscopy	Arthroscopic Meniscectomy	<div></div>						<div> OMEROS</div>
MASP (OMS721) - Complement-Mediated Disorders	TMA (aHUS, TTP, Transplant-Related TMA)	<div></div>						
PDE10 (OMS824) - CNS Disorders	Schizophrenia	<div></div>						
PDE10 (OMS824) - CNS Disorders	Huntington's Disease	<div></div>						
PPARγ (OMS405) - Addiction	Opioid and Nicotine Addiction	<div></div>						
OMS201 - Urology	Ureteroscopy	<div></div>						
Preclinical Programs								
PDE7 (OMS527) - CNS Disorders	Addictions and Compulsive Disorders; Movement Disorders	<div></div>						<div> OMEROS</div>
Plasmin (OMS616) - Bleeding Disorders	Surgical and Traumatic Bleeding	<div></div>						
MASP (OMS906) - Alternative Pathway Disorders	Paroxysmal Nocturnal Hemoglobinuria	<div></div>						
GPR17 - CNS Disorders	Demyelinating Disorders	<div></div>						
GPCR Platform	Multiple Disorders Across Therapeutic Areas	<div></div>						
Antibody Platform	Multiple Disorders Across Therapeutic Areas	<div></div>						



Next-Generation Therapies Transforming Patient Care Today