

# AXSOME

## THERAPEUTICS

May 2018

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# Developing novel therapies for CNS disorders.

Axsome is addressing growing markets, where current treatment options are limited or inadequate, by leveraging well-characterized compounds to create novel therapeutics to meet unmet medical needs and improve the lives of patients.

# Our Technologies

Axsome's medicinal chemistry and formulation technologies enable new and innovative medicines to treat CNS conditions.



# Our Candidates and Pipeline

- Five differentiated clinical-stage assets targeting significant and growing markets.
- Patent protection to 2034, worldwide rights.

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-05 (DM + BUP)	Treatment Resistant Depression: Fast Track Granted			Ongoing
	Agitation in Alzheimer's Disease: Fast Track Granted			Ongoing
	Smoking Cessation			Ongoing
AXS-09 (DM + S-BUP)	CNS Disorders			
AXS-02 (DZT)	Knee OA with BMLs: SPA Received; Fast Track Granted			Ongoing
	CLBP with MCs			
AXS-07 (MoSEIC™ Mx + Riz)	Migraine			
AXS-06 (MoSEIC™ Mx + Eso)	OA and RA			

Abbreviations: BML = Bone Marrow Lesions; BUP = Bupropion; CLBP = Chronic Low Back Pain; DM = Dextromethorphan; DZT = Disodium Zoledronate Tetrahydrate; Eso = Esomeprazole; MC = Modic Changes; Mx = Meloxicam; OA = Osteoarthritis; RA = Rheumatoid Arthritis; Riz = Rizatriptan; S-BUP = Esbupropion; SPA = Special Protocol Assessment.

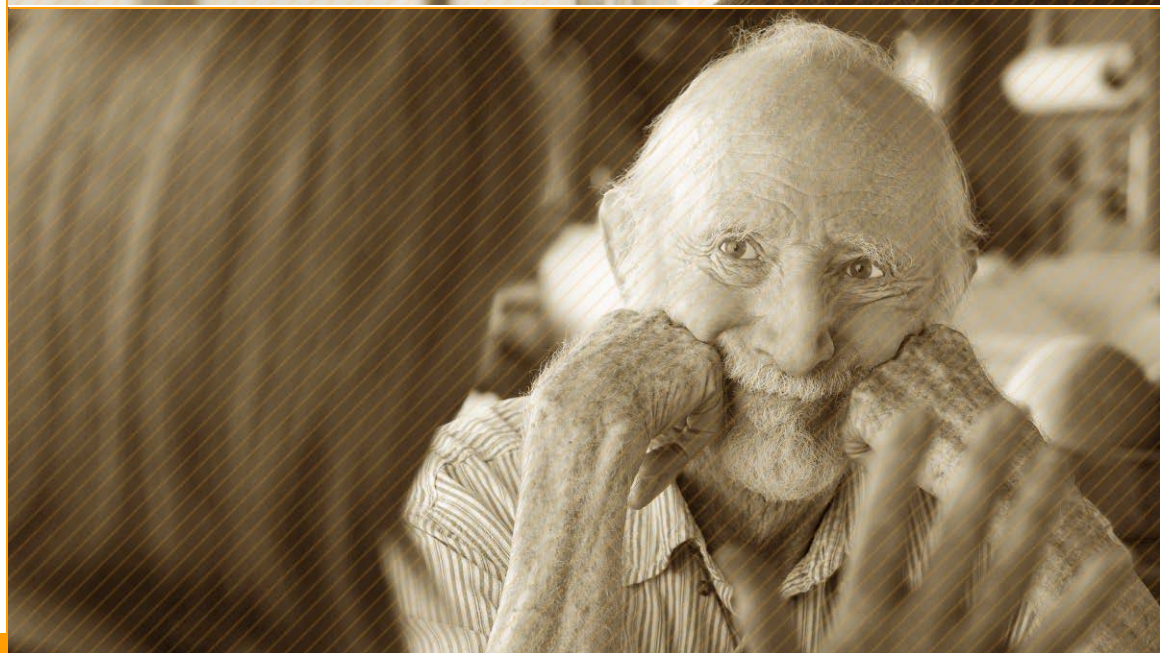


# AXS-05

## Dextromethorphan (DM) + Bupropion (BUP)

Novel therapy for CNS disorders:

- Treatment Resistant Depression (TRD)
- Agitation in Alzheimer's Disease (AD)
- Smoking Cessation



# CNS Disorders: Mechanisms of Action

## Pharmacodynamic Synergy

Mechanism of Action	AXS-05		
	DM	BUP	DM+BUP
NMDA Receptor Antagonist	✓		✓
Sigma-1R Agonist	✓		✓
Norepinephrine Reuptake Inhibitor	✓	✓	✓
Serotonin Reuptake Inhibitor	✓		✓
Dopamine Reuptake Inhibitor		✓	✓
Nicotinic ACh Receptor Antagonist	✓	✓	✓

DM = Dextromethorphan; BUP = Bupropion.

✓ Present

# CNS Disorders:

## Mechanisms of Action and Relevant Indications

Mechanism of Action	Pharmacodynamic Synergy			Relevant Indications <sup>1</sup>								Related Agents <sup>2</sup>
	DM	BUP	AXS-05 DM+BUP	ADHD	Anxiety	Alzheimer's	Depression	Fibromyalgia	OCD	Pain	Smoking cessation	
NMDA Receptor Antagonist	✓		✓									<ul style="list-style-type: none"> <li>• Ketamine</li> <li>• Memantine (Namenda®)</li> </ul>
Sigma-1R Agonist	✓		✓									<ul style="list-style-type: none"> <li>• Fluvoxamine (Luvox®)</li> <li>• Donepezil (Aricept®)</li> </ul>
Norepinephrine Reuptake Inhibitor	✓	✓	✓									<ul style="list-style-type: none"> <li>• Duloxetine (Cymbalta®)</li> <li>• Venlafaxine (Effexor®)</li> </ul>
Serotonin Reuptake Inhibitor	✓		✓									<ul style="list-style-type: none"> <li>• Escitalopram (Lexapro®)</li> <li>• Fluoxetine (Prozac®)</li> <li>• Sertraline (Zoloft®)</li> </ul>
Dopamine Reuptake Inhibitor		✓	✓									<ul style="list-style-type: none"> <li>• Bupropion (Wellbutrin®)</li> </ul>
Nicotinic ACh Receptor Antagonist	✓	✓	✓									<ul style="list-style-type: none"> <li>• Bupropion (Wellbutrin®)</li> </ul>

DM = Dextromethorphan; BUP = Bupropion.

✓ Present

■ Relevant

1. Indications listed are associated with the mechanism of action and are not related to either DM or BUP, unless specifically noted.

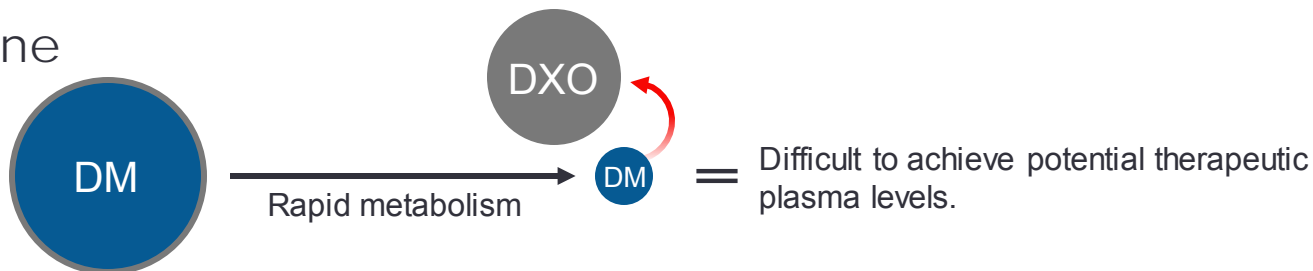
2. Agents do not contain DM or BUP, unless specifically noted.



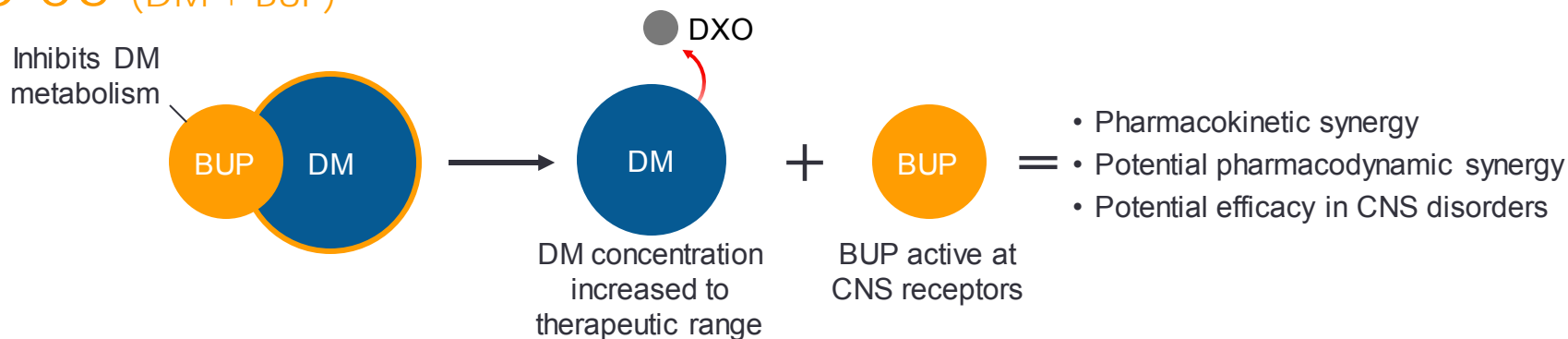
# CNS Disorders:

## Novel Therapy for CNS Disorders

DM Alone



AXS-05 (DM + BUP)



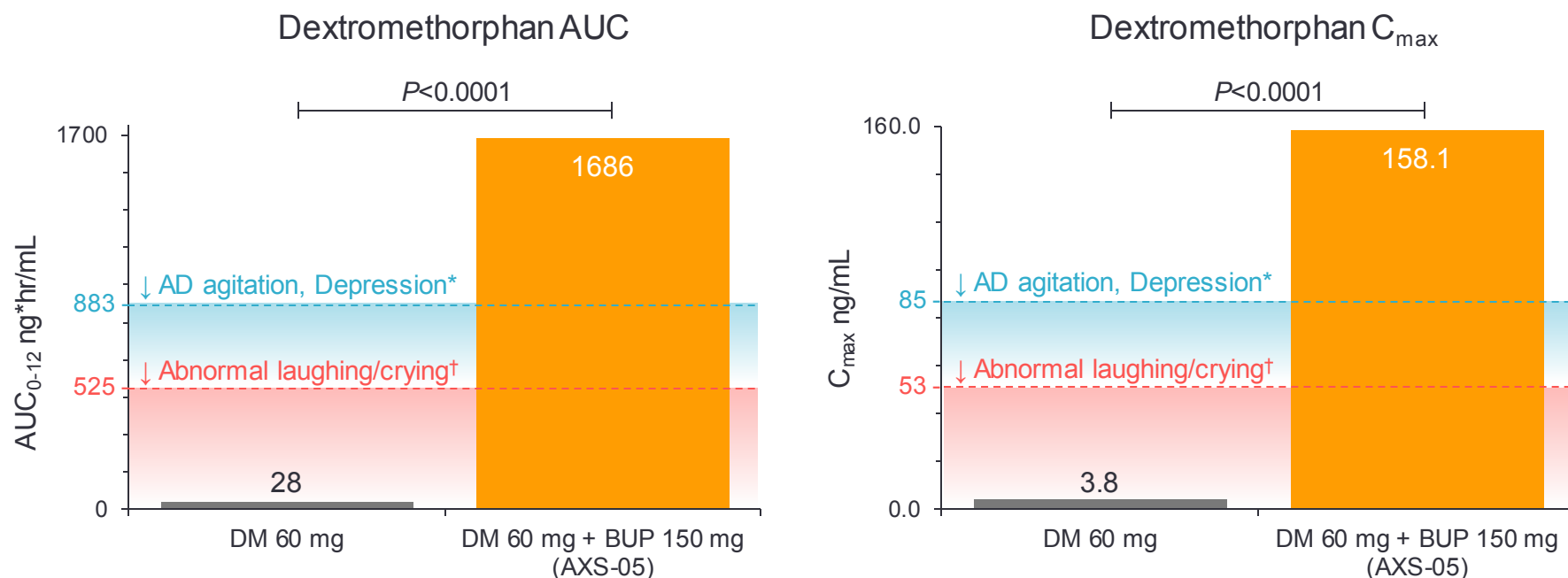
DM = Dextromethorphan; DXO = Dextrorphan; BUP = Bupropion.

- Phase 1 trials with AXS-05 completed:
  - Significant increase in DM plasma levels.
- Phase 3 trials in TRD and AD Agitation initiated.

### IP Overview

- 25 issued patents – protection through 2034.

# CNS Disorders: Phase 1 Results



DM concentrations associated with reported therapeutic responses shown (dotted lines).

\* DM plasma concentrations reported with dose (DM 30 mg + Q 10 mg) resulting in reduction of agitation symptoms in AD patients, and of depressive symptoms in AD and PBA patients.

† DM plasma concentrations reported with dose (DM 20 mg + Q 10 mg) resulting in reduction in emotional symptoms in PBA patients.

Axsome data on file.

Therapeutic DM concentrations from NDA 021879, FDA Clinical Pharmacology Review.

DM, Dextromethorphan; Q, Quinidine; BUP, Bupropion; AD, Alzheimer's disease; PBA, pseudobulbar affect

# CNS Disorders:

## TRD Overview

- Major Depressive Disorder (MDD) is a leading cause of disease burden in the US.<sup>4</sup>
- 63% and 44% of MDD patients have inadequate response to initial therapy and second line therapy, respectively.<sup>2</sup>
- Only 1 approved drug for TRD = unmet medical need.
- AXS-05 combines the MOA of 4 distinct anti-depressant drug classes into one novel oral therapeutic.
- DM antidepressant effects demonstrated preclinically and clinically.
- Phase 3 interim futility analysis: IDMC recommended trial continuation.



3M patients  
in the U.S.<sup>1-3</sup>

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-05 (DM + BUP)	Treatment Resistant Depression: Fast Track Granted			Ongoing

Abbreviations: DM = Dextromethorphan; BUP = Bupropion.

1. Marcus SC, Olsson M. *Arch Gen Psychiatry* 2010;67:1265-1273.

2. Rush AJ, et al. *Am J Psychiatry* 2006;163:1905-1917.

3. U.S. Census Bureau, Population April 1, 2010 to July 1, 2013.

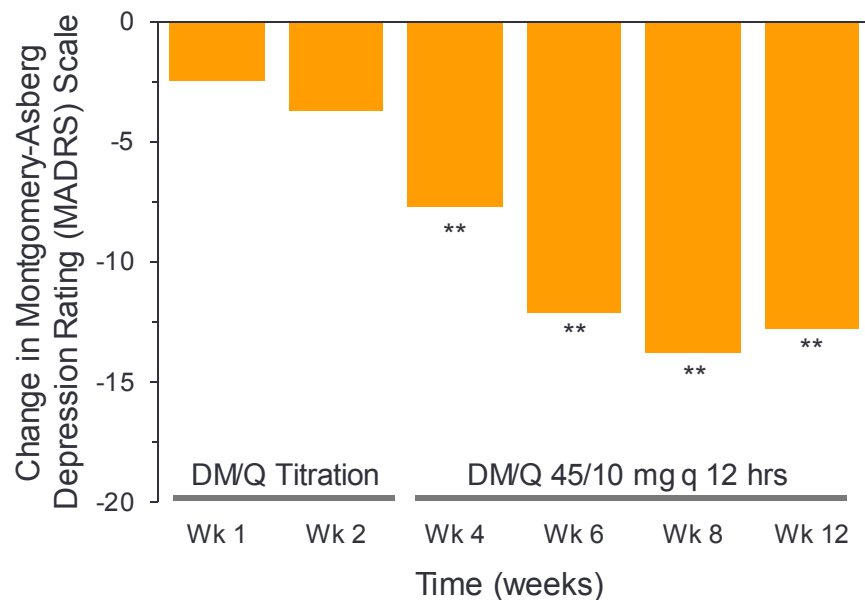
4. Mathers CD, *PLoS Med* 2006; 3(11): e442.

# CNS Disorders:

## TRD Clinical Rationale

- DM and metabolic inhibitor reduce depressive symptoms in TRD and in AD.

Symptom Reduction in TRD Patients  
Treated with DM and Metabolic Inhibitor<sup>1</sup>



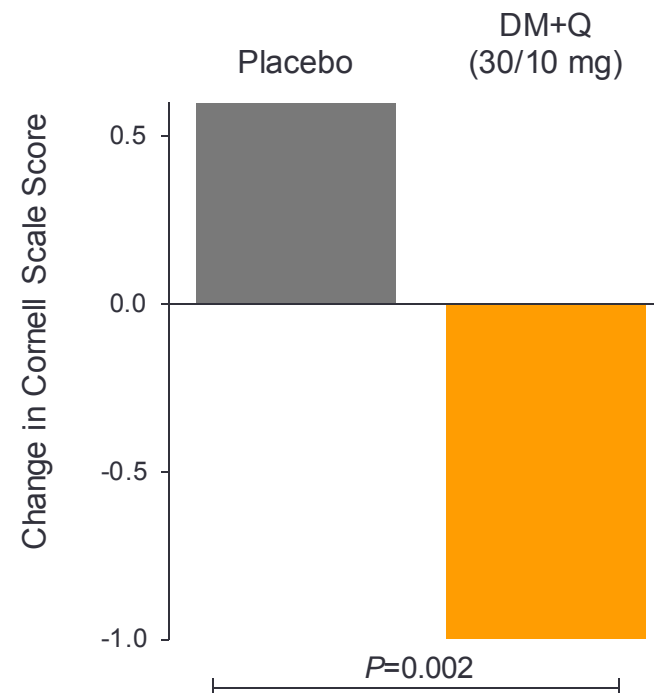
- Failed 2 to 10 prior treatments
- 45% of patients had  $\geq 50\%$  reduction in MADRS

\*\*  $P < 0.01$  versus baseline

1. Murrough J, et al. *J Affect Disord.* 2017;218:277-283.

2. Cummings J, et al. *JAMA.* 2015;314:1242-1254.

Depressive Symptom Reduction in AD Agitation  
Patients Treated with DM and Metabolic Inhibitor<sup>2</sup>

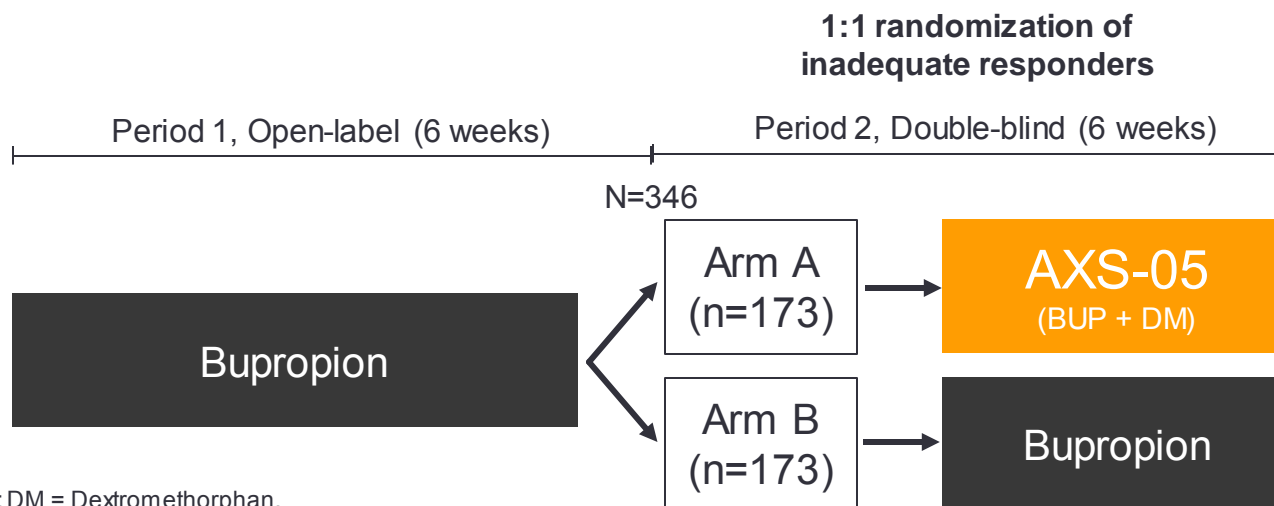


# CNS Disorders:

## TRD Phase 3 Design



A Phase 3 trial to assess the efficacy and safety of  
**AXS-05** in the treatment of TRD.



BUP = Bupropion; DM = Dextromethorphan.

- **Primary Endpoint:** Change in depression score from randomization to end of study, measured using the Montgomery-Asberg Depression Rating Scale (MADRS).
- **Key Inclusion Criteria:**
  - Male or female 18-65 years old
  - History of inadequate response to 1 or 2 adequate antidepressant treatments
- **Interim futility analysis:** Conducted at approximately 40% target randomized subjects. IDMC recommended trial continuation.
- **Interim efficacy analysis:** Planned at approximately 60% target randomized subjects.



# CNS Disorders:

## Agitation in AD Overview

- Agitation and aggression seen in approximately 45% of AD patients during 5-year period.<sup>3</sup>
- Characterized by emotional distress, aggressive behaviors, disruptive irritability, disinhibition, and caregiver burden.<sup>4</sup>
- Associated with<sup>4,5</sup>:
  - Accelerated cognitive decline
  - Earlier nursing home placement
  - Increased mortality
- No approved medication = unmet medical need.
- Proof of concept: DM plus metabolic inhibitor reduced agitation in AD patients.
- Phase 2/3 ongoing.



2M patients  
in the U.S.<sup>1,2</sup>

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-05 (DM + BUP)	Agitation in Alzheimer's Disease: Fast Track Granted			Ongoing

Abbreviations: DM = Dextromethorphan; BUP = Bupropion.

1. Ryu, SH, et al. *Am J Geriatr Psychiatry*. 2005;13:976-983.

2. Hebert, LE, et al. *Neurology*. 2013;80:1778-1783.

3. Steinberg M, et al. *Int J Geriatr Psychiatry*. 2008;2:170-177.

4. Antonsdottir IM, et al. *Expert Opin Pharmacother*. 2015;11:1649-1656.

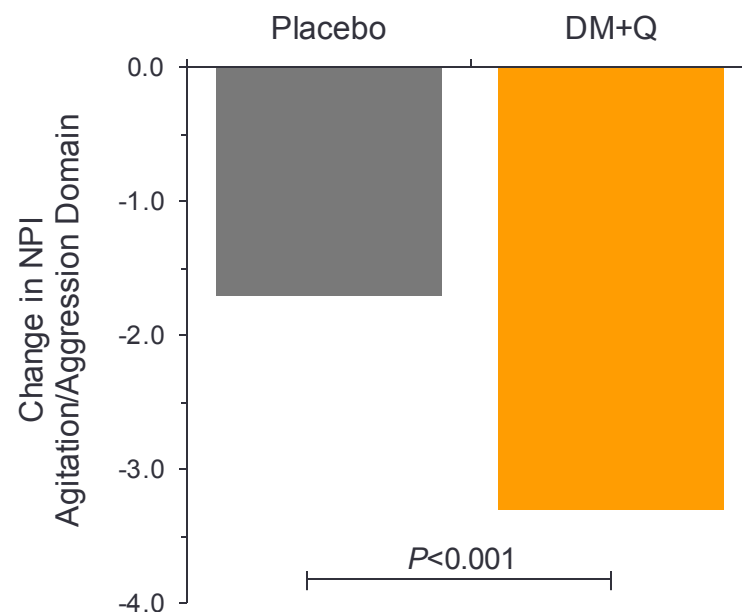
5. Rabins PV et al. *Alzheimers Dement*. 2013; 9:204-207.

# CNS Disorders:

## Agitation in AD Clinical Rationale

- Randomized, double-blind, placebo-controlled, two-stage trial.
  - Placebo (n=125), 30 mg DM + 10 mg quinidine (Q) (n=93), for stage 1.
- DM+Q treatment reduced agitation/aggression in AD by 46% vs. 24% for placebo ( $P<0.001$ )—primary endpoint.
- Statistically significant improvement in multiple secondary endpoints.
- DM plasma levels achieved with AXS-05 in target therapeutic range.
- Potential for additional contribution from bupropion component of AXS-05.

Change in Agitation/Aggression Scores in AD with DM and Metabolic Inhibitor Quinidine (Q)



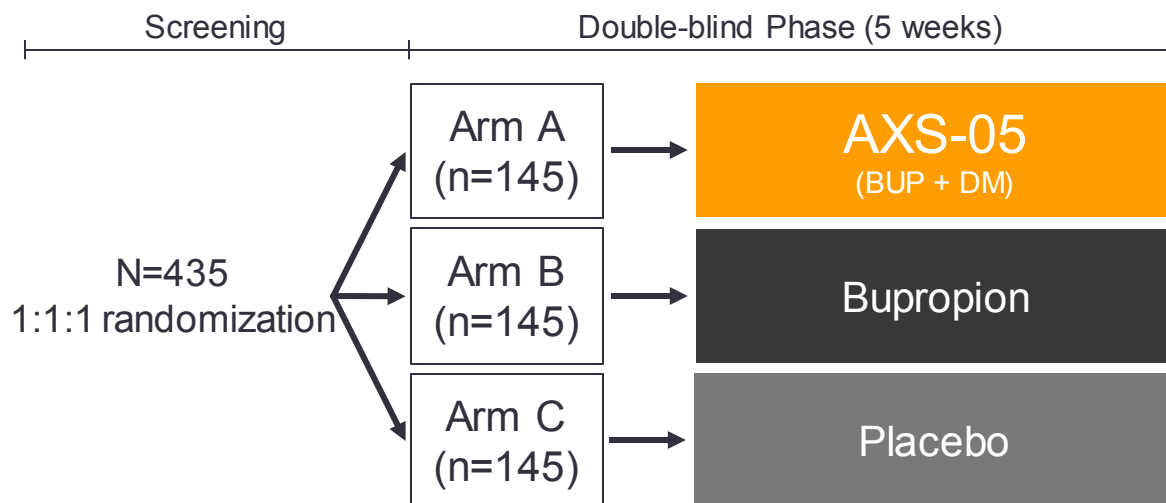
Cummings J, et al. *JAMA*. 2015;314:1242-1254.

# CNS Disorders:

## Agitation in AD Phase 2/3 Design

**ADVANCE**  
STUDY

A Phase 2/3 trial to assess the efficacy and safety of  
**AXS-05** in the treatment of Agitation in AD.



BUP = Bupropion; DM = Dextromethorphan.

- **Primary Endpoint:** Cohen-Mansfield Agitation Inventory (CMAI).
- **Key Inclusion Criteria:**
  - Diagnosis of probable Alzheimer's disease
  - Clinically significant agitation
- Interim futility analysis: Planned at approximately 30% target randomized subjects.
- Interim efficacy analysis: Planned at approximately 60% target randomized subjects.

# CNS Disorders:

## Smoking Cessation Overview

- Smoking is single largest cause of preventable death in the U.S.<sup>1</sup>
- 70% of smokers want to quit and only 3-5% who attempt to quit without assistance are successful for 6-12 months.<sup>2</sup>
- DM component of AXS-05 significantly reduced nicotine self-administration in nicotine-dependent rats.
- Bupropion component of AXS-05 has been found to be effective for smoking cessation in clinical trials.
- Axsome entered into a research collaboration with Duke University to evaluate AXS-05 in a Phase 2 clinical trial in smokers attempting to quit.
- Phase 2 controlled trial initiation anticipated in 1H 2018.



40M patients  
in the U.S.<sup>1</sup>

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-05 (DM + BUP)	Smoking Cessation			Ongoing

Abbreviations: DM = Dexamethorphan; BUP = Bupropion.

1. U.S. Department of Health and Human Services. The Health Consequences of Smoking: 50 Years of Progress. A Report of the Surgeon General. 2014.

2. Hughes JR, et al. *Addiction*. 2004;99(1):29-38.

# CNS Disorders:

## AXS-09 (Esbupropion + DM) Overview

- Esbupropion and DM fixed-dose combination
- Esbupropion is the chirally pure *S*-enantiomer of bupropion.
- Phase 1 trial completed:
  - Pharmacokinetic trial of AXS-09, *R*-bupropion and dextromethorphan, single-entity *S*-bupropion, or single-entity *R*-bupropion
  - Substantial increases in DM plasma concentrations with AXS-09 ( $p < 0.0001$  day 1 versus day 8)
  - DM concentrations with AXS-09 comparable to AXS-05
  - AXS-09 was well tolerated
- To be developed in future CNS indications



Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-09 (DM + S-BUP)	CNS Disorders			

Abbreviations: DM = Dextromethorphan; S-BUP = Esbupropion.



# AXS-02

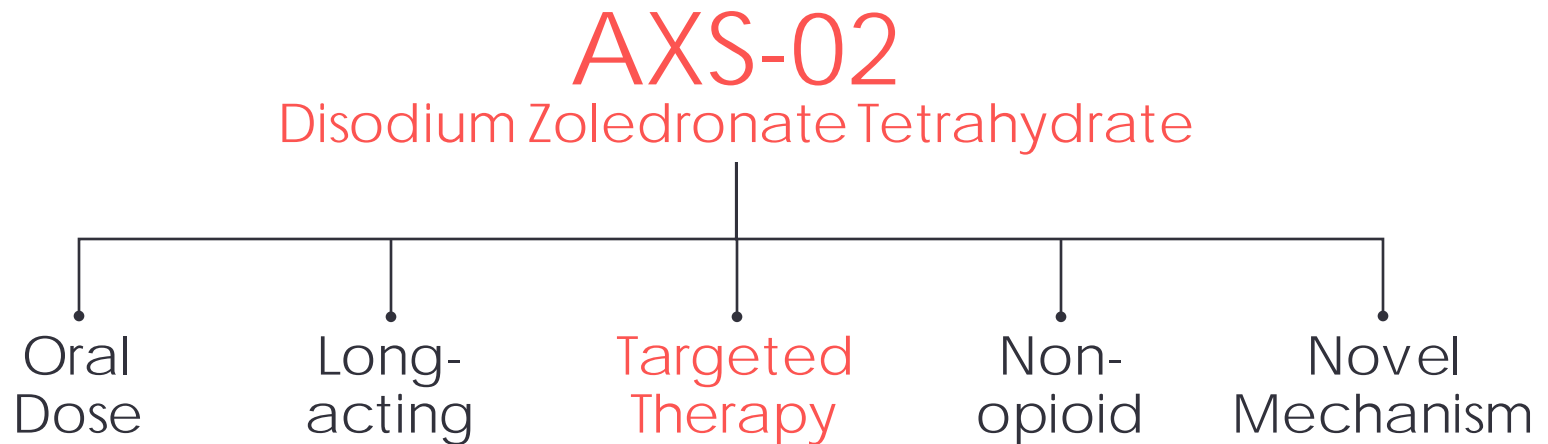
## Disodium Zoledronate Tetrahydrate

Novel therapy for chronic pain:

- Knee Osteoarthritis (OA) with Bone Marrow Lesions (BMLs)
- Chronic Low Back Pain (CLBP) with Modic Changes (MCs)



# Chronic Pain: Differentiated Therapy



## Mechanisms of Action



Inhibits  
bone-resorbing  
osteoclasts



Downregulates  
acid-sensing<sup>†</sup>  
ion channels



Reduces  
pro-inflammatory  
cytokine production



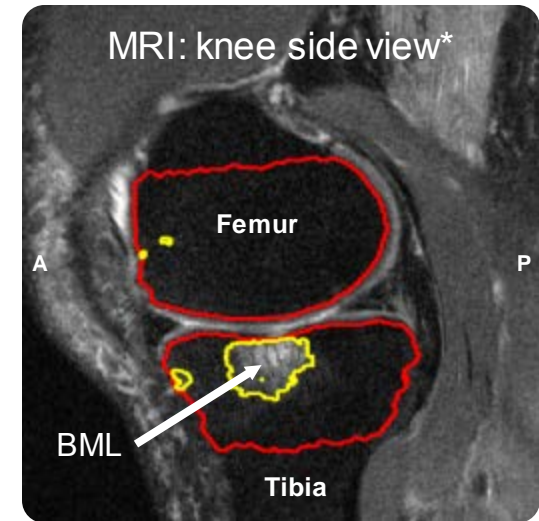
Anti-angiogenic

<sup>†</sup>Acid is a well known cause of pain.

# Chronic Pain:

## Knee OA with BMLs Overview

- Bone marrow lesions (BMLs) on MRI are associated with pain in knee osteoarthritis (OA).<sup>1</sup>
- BMLs are regions of increased bone turnover, and reduced mineral density.<sup>2,3</sup>
- Zoledronic acid inhibits bone resorption and increases mineral density.
- Phase 3 trial initiated based on positive Phase 2 results with IV zoledronic acid.
- Phase 3 interim analysis: IDMC recommended continuation to full enrollment



7M patients  
in the U.S.<sup>4-9</sup>

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-02 (DZT)	Knee OA with BMLs: SPA Received; Fast Track Granted			Initiated

Abbreviations: DZT = Disodium Zoledronate Tetrahydrate.

\* MRI showing BML in medial tibia from Driban, et al. *Arthritis Res Ther.* 2013;15:R112.

1. Driban JB, et al. *Arthritis Res Ther.* 2013;15:R112.

2. Hunter DJ, et al. *Arthritis Res Ther.* 2009;11:R11.

3. Kazakia GJ, et al. *Osteoarthritis Cartilage.* 2013;21:94-101.

4. Lawrence RC, et al. *Arthritis Rheum.* 2008;58:26-35.

5. Zhang Y, Jordan. *JM Clin Geriatr Med.* 2010;26:355-69.

6. Tanamas SK, et al. *Rheumatology.* 2010;49:2413-19.

7. Guermazi A, et al. *BMJ.* 2012;345:e5339.

8. Jensen OK, et al. *Spine J.* Feb. 14, 2014;pii:S1529-9430(14)00214-9.

9. U.S. Census Bureau, Population April 1, 2010 to July 1, 2013.

# Chronic Pain:

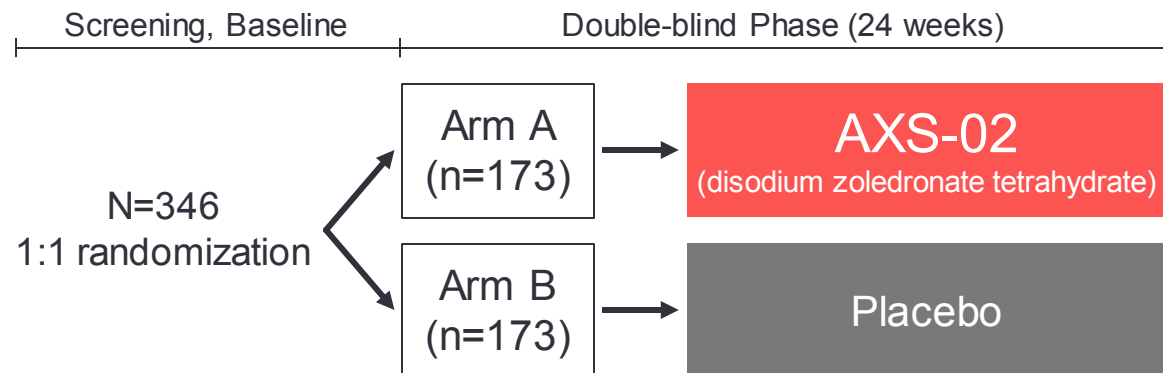
## Knee OA with BMLs Phase 3 Design

**coast-1**

Clinical Knee OA Symptom  
Treatment 1 Study

Special Protocol  
Assessment (SPA)  
received

A Phase 3 trial to assess the efficacy and safety of  
**AXS-02** in the treatment of pain of knee OA associated with BMLs.



- **Primary Endpoint:** Change in pain intensity from baseline to week 24, measured using the 0-10 Numerical Rating Scale (NRS).
- **Key Inclusion Criteria:**
  - Male at least 50 years of age or postmenopausal female, with knee OA and BMLs
  - Moderate or worse knee pain
- **Dosage:** Once per week for six weeks; no drug for remainder of double-blind phase.

# Chronic Pain:

## CLBP with MCs Overview

- Modic changes (MCs) type 1 (M1) on MRI are associated with chronic low back pain (CLBP).<sup>1</sup>
- Increased bone turnover on bone scan is seen in M1 lesions.<sup>2</sup>
- Increased pro-inflammatory cytokines, and vascular density seen in M1 lesions.<sup>3</sup>
- Zoledronic acid reduces bone turnover, suppresses the production of inflammatory mediators, and is anti-angiogenic.
- Phase 2 results: Zoledronic acid reduced pain in patients with CLBP.
- FDA clearance received for IND for Phase 3 trial – initiation planned following readouts from CREATE-1 and STRIDE-1.
- Issued U.S. patents: protection into 2034 – uses of oral zoledronic acid for low back pain.



1.6M patients  
in the U.S.<sup>4-7</sup>

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-02 (DZT)	CLBP with MCs			

Abbreviations: DZT = Disodium Zoledronate Tetrahydrate.

\* MRI showing modic type 1 lesions from Luoma K, et al. *European Congress of Radiology (ECR)*. 2014;Poster B-0458.

1. Zhang Y, et al. *Eur Spine J*. 2008;17:1289-1299.

2. Järvinen J, et al. *Spine: ISSLS Society Meeting Abstracts*. Oct. 2011;Volume Suppl, Abstract GP127.

3. Rahme R, Moussa R. *Am J Neuroradiol*. 2008;29:838–42.

4. Lawrence RC, et al. *Arthritis Rheum*. 2008;58:26-35.

5. Zhang Y, Jordan. *JM Clin Geriatr Med*. 2010;26:355–69.

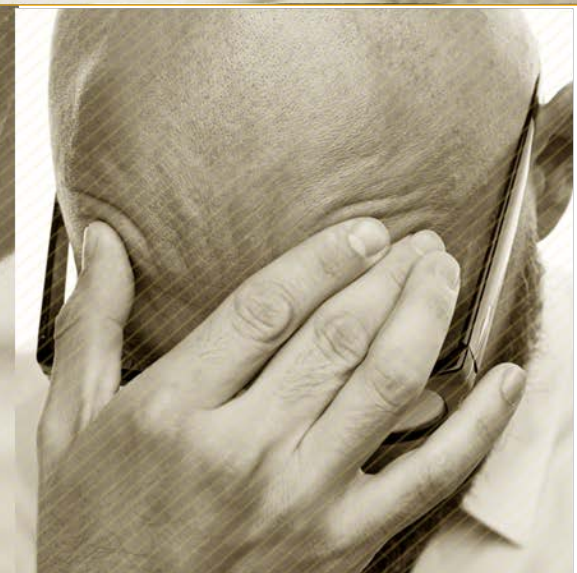
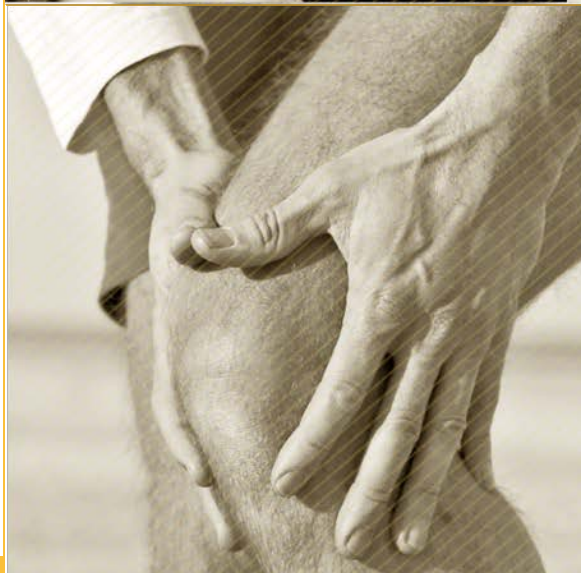
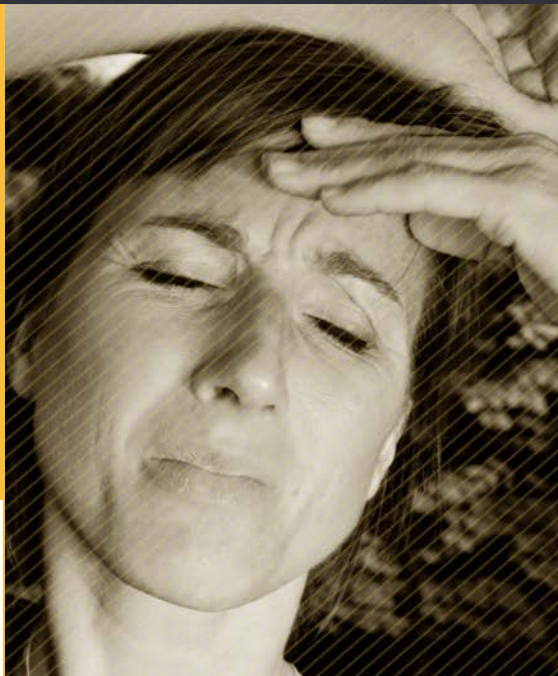
6. Jensen OK, et al. *Spine J*. Feb. 14, 2014;pii:S1529-9430(14)00214-9. 7. U.S. Census Bureau, Population April 1, 2010 to July 1, 2013.



# MoSEIC™ Meloxicam

Novel therapies:

- AXS-07 – Migraine
- AXS-06 – OA and RA



# Migraine, OA and RA:

## MoSEIC™ Meloxicam Overview

- MoSEIC meloxicam is a potent, oral, rapidly-absorbed, once-daily, non-opioid, COX-2 preferential, pain therapeutic.
- Standard meloxicam has an extended  $T_{\max}$  (4-6 hours) which delays its onset of action.<sup>1,2</sup>
- Axsome's MoSEIC (Molecular Solubility Enhanced Inclusion Complex) technology substantially increases the rate of absorption of meloxicam while maintaining its approximately 20-hour half-life.
- Phase 1 results: 9 times faster  $T_{\max}$ , higher  $C_{\max}$  and similar half-life, compared to Mobic®.
- Potential utility for migraine, and the signs and symptoms of OA and RA.
- AXS-07 is a fixed-dose combination of MoSEIC meloxicam and rizatriptan.
- AXS-06 is a fixed-dose combination of MoSEIC meloxicam and esomeprazole (to reduce risk of NSAID-associated ulcers).

### IP Overview

- 1 issued patent – protection through 2036.
- Pharmacokinetic patents
- More than 10 U.S. and international applications.

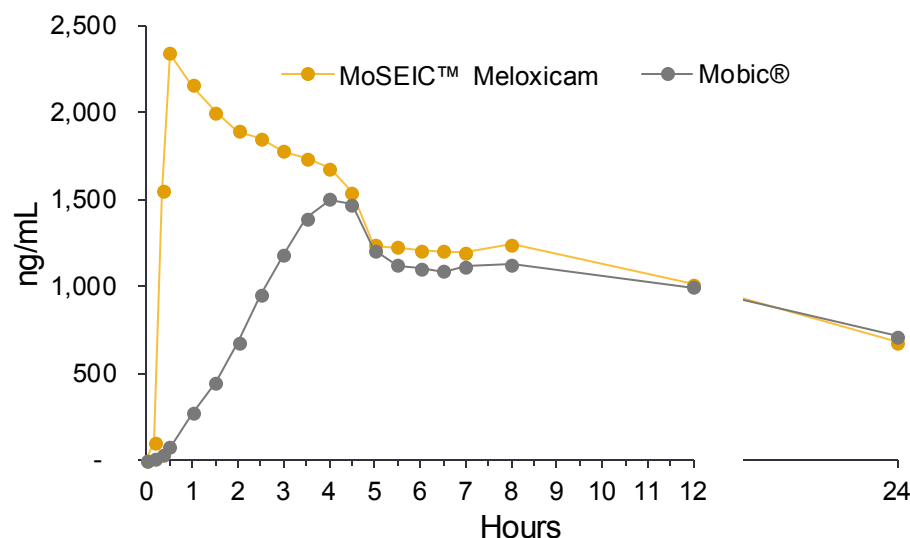
1. Mobic® (meloxicam) FDA Package Insert.

2. Euler-Ziegler et al., *Inflamm Res* 50, Supplement 1 (2001) S5–S9.

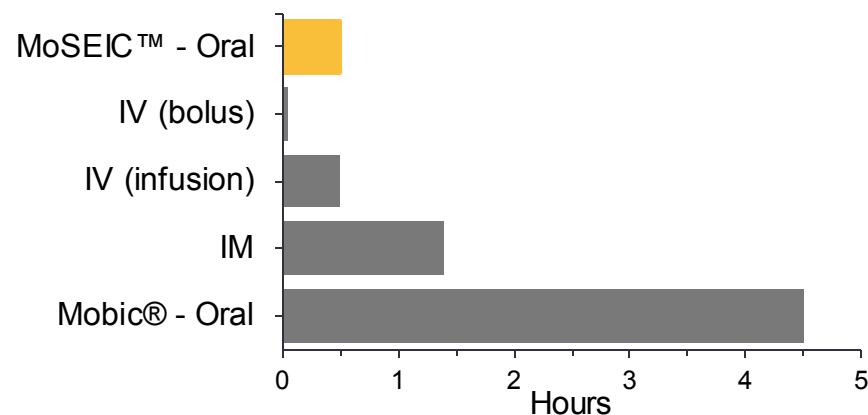
# Migraine, OA and RA:

## MoSEIC™ Meloxicam Phase 1 Results

Mean Meloxicam Concentrations



Meloxicam  $T_{max}$  after 15 mg Dose

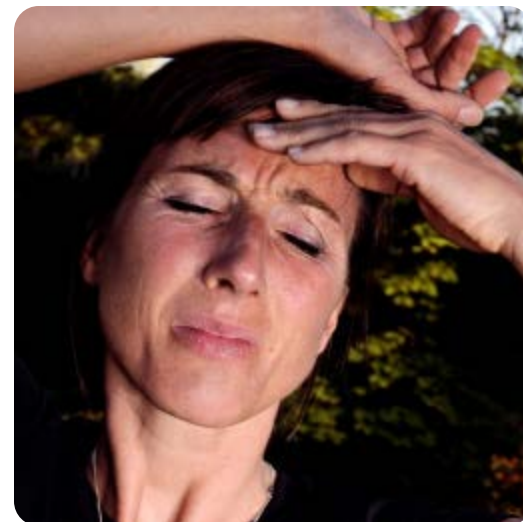


- MoSEIC meloxicam  $T_{max}$  9 times faster than Mobic® (0.5 hour versus 4.5 hours, respectively,  $p < 0.0001$ ).
- Therapeutic plasma levels achieved within 15 minutes of oral dosing of MoSEIC meloxicam.
- MoSEIC meloxicam had higher mean  $C_{max}$  ( $p = 0.0018$ ), faster time to therapeutic plasma concentration ( $p < 0.0001$ ), and time to half-maximal plasma concentration ( $p < 0.0001$ ) as compared to Mobic®.
- Terminal half-lives were approximately 20 hours for MoSEIC meloxicam and 22 hours for Mobic®.

Sources: Axsome data on file. IV and IM data from Euler-Ziegler et al., *Inflamm Res* 50, Supplement 1 (2001) S5–S9.

# AXS-07: MoSEIC™ Meloxicam + Rizatriptan for Migraine

- Meloxicam is a new molecule for migraine—not currently approved or used for this indication due to prolonged  $T_{\max}$
- MoSEIC delivery enables its use in abortive treatment of migraine
  - Rapid  $T_{\max}$  of MoSEIC meloxicam is ideal for migraine treatment
  - Extended half-life of MoSEIC meloxicam should lead to lower symptom recurrence
- AXS-07 combines unique PK of MoSEIC meloxicam with proven efficacy of rizatriptan
- FDA Pre-IND written guidance received
- Phase 3 initiation anticipated in 2018



37M patients  
in the U.S.<sup>1</sup>

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-07 (MoSEIC™ Mx + Riz)	Migraine			

Abbreviations: Mx = Meloxicam; Riz = Rizatriptan.

1. Fleis JR, et al., *Summary health statistics for U.S. adults: National Health Interview Survey, 2009. National Center for Health Statistics. Vital Health Stat 10(249). 2010.*

# AXS-07:

## Differentiated Clinical Profile for Migraine



Rapid absorption  
& onset of action

Based on rapid absorption of MoSEIC meloxicam and expected additive effect of AXS-07 components



Strong & consistent  
pain relief

Potential for superior efficacy as compared to current treatments based on expected additive effect of AXS-07 components



Sustained  
pain relief

Based on extended MoSEIC meloxicam half-life and expected additive effect of AXS-07 components



Pharmaco-  
economic benefits

Potentially superior efficacy expected to result in reduced use of medication and medical services, reduced absenteeism and loss of productivity



# AXS-06:

## MoSEIC™ Meloxicam + Esomeprazole for OA & RA

- AXS-06 is a fixed-dose combination of MoSEIC™ meloxicam and esomeprazole
- Being developed to treat OA and RA, and to reduce the risk of NSAID-associated upper GI ulcers
- Potentially best-in-class NSAID profile:
  - Oral administration with IV-like onset of action
  - Long half-life for sustained effect and once-daily dosing
  - Improved GI safety from esomeprazole component
- Positive Phase 1 results: therapeutic meloxicam concentrations within 15 mins, gastroprotective esomeprazole concentrations
- FDA Pre-IND written guidance received
- AXS-06 is Phase 3-ready

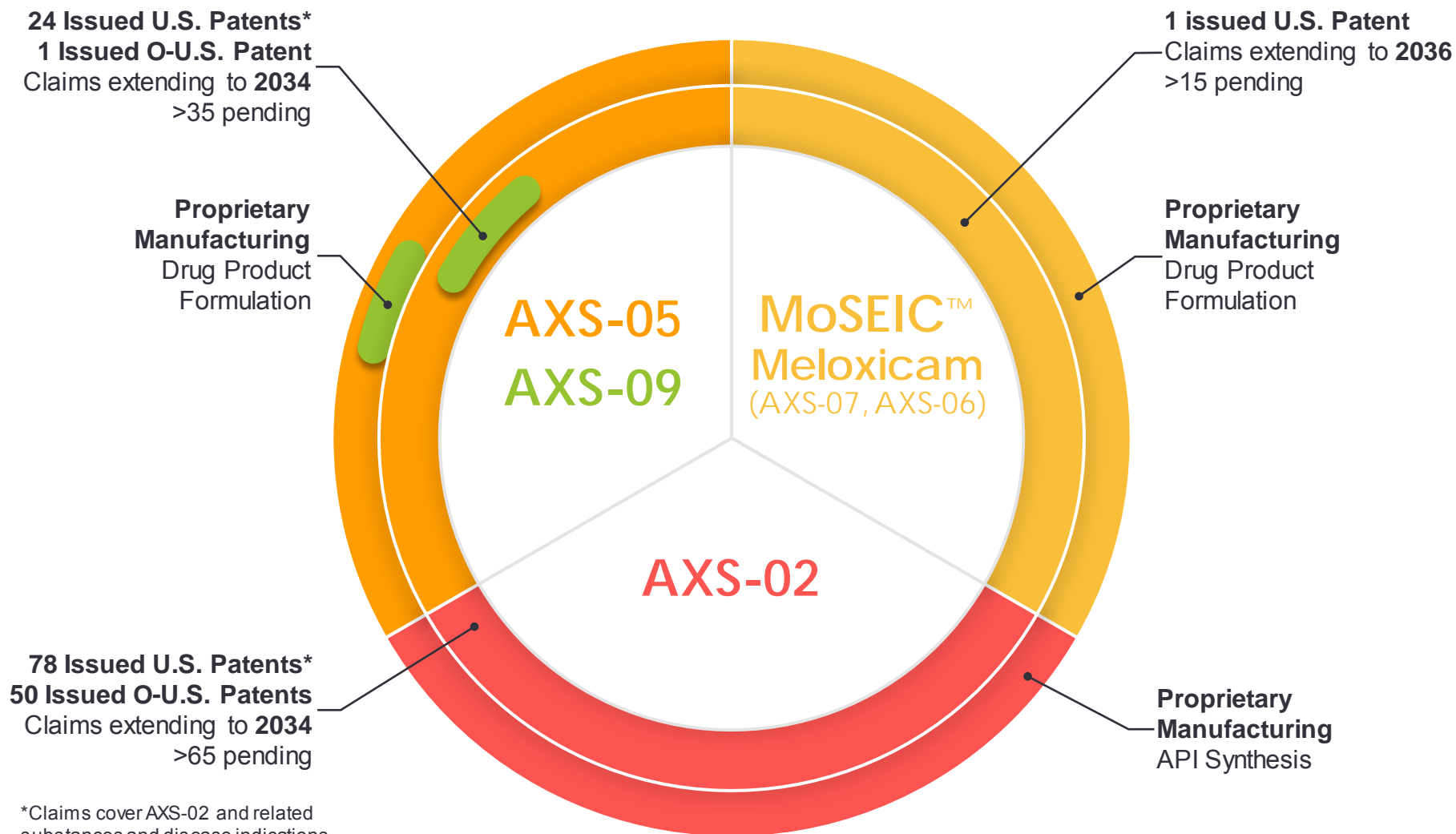


**120M** NSAID TRx  
per year  
in the U.S.

Product Candidate	Preclinical	Phase 1	Phase 2	Phase 3
AXS-06 (MoSEIC™ Mx + Eso)	OA and RA			Phase 3 ready

Abbreviations: Eso = Esomeprazole; Mx = Meloxicam; OA = Osteoarthritis; RA = Rheumatoid Arthritis.

# Barriers to Entry



# Our Team

## Management

Herriot Tabuteau, MD  
Founder & CEO

Nick Pizzie, CPA, MBA  
CFO

Cedric O'Gorman, MD, MBA  
SVP, Clinical Development & Medical Affairs

Mark Jacobson, MA  
SVP, Operations

Robert Niecestro, PhD  
VP, Clinical & Regulatory



AXSOME THERAPEUTICS

## Board of Directors

Roger Jeffs, PhD  
Former President, Co-CEO, Director  
**United Therapeutics Corp.**  
Prior positions at Amgen and Burroughs Wellcome

Myrtle Potter  
Former President, COO  
**Genentech**  
Prior positions at Bristol-Myers Squibb and Merck

Mark Saad  
Former CFO  
**Bird Rock Bio, Inc.**  
Former COO of the Global Healthcare Group at UBS

Mark Coleman, MD  
Medical Director  
**National Spine and Pain Centers**  
Diplomat of the American Board of Anesthesiology

Herriot Tabuteau, MD  
Chairman

# Key Financial Information

As of March 31, 2018	
Cash:	\$26.6 Million
Debt (Face Value) <sup>1</sup> :	\$8.9 Million
Common Shares Outstanding:	25.5 Million
Options and Warrants Outstanding <sup>2</sup> :	4.9 Million

- **Financial guidance:** Cash anticipated to fund operating requirements into the third quarter of 2019.

1. Book value of \$9.2 million.

2. Consists of 2.8 million options and 2.1 million warrants.

# Clinical Milestones

Product Candidate	Indication	2018	2019
AXS-05 (DM + BUP)	<b>TRD</b>	<ul style="list-style-type: none"> <li>✓ <b>STRIDE-1</b> interim analysis</li> <li>● <b>STRIDE-1</b> interim efficacy analysis (2H 2018)</li> <li>● <b>STRIDE-1</b> top-line results (2H 2018/1H 2019)</li> </ul>	
	<b>AD Agitation</b>	<ul style="list-style-type: none"> <li>● <b>ADVANCE-1</b> interim analysis (2H 2018)</li> </ul>	<ul style="list-style-type: none"> <li>● <b>ADVANCE-1</b> interim efficacy analysis</li> <li>● <b>ADVANCE-1</b> top-line results (2H 2019/1H 2020)</li> </ul>
	<b>Smoking Cessation</b>	<ul style="list-style-type: none"> <li>✓ <b>Ph 2</b> trial start</li> </ul>	<ul style="list-style-type: none"> <li>● <b>Ph 2</b> top-line results</li> </ul>
AXS-09 (DM + S-BUP)	<b>CNS Disorders</b>	<ul style="list-style-type: none"> <li>✓ <b>Ph 1</b> trial results</li> </ul>	
AXS-07 (MoSEIC™ Mx + Riz)	<b>Migraine</b>	<ul style="list-style-type: none"> <li>● <b>Ph 3</b> trial start</li> </ul>	<ul style="list-style-type: none"> <li>● <b>Ph 3</b> top-line results</li> </ul>
AXS-02 (DZT)	<b>Knee OA</b>	<ul style="list-style-type: none"> <li>✓ <b>COAST-1</b> interim analysis</li> </ul>	

Abbreviations: AD= Alzheimer's Disease; BUP = Bupropion; CLBP = Chronic Low Back Pain; DM = Dextromethorphan; DZT = Disodium Zoledronate Tetrahydrate; Eso = Esomeprazole; ; Mx = Meloxicam; OA = Osteoarthritis; RA = Rheumatoid Arthritis; Riz = Rizatriptan; S-BUP = Esbupropion; TRD = Treatment Resistant Depression.

✓ Accomplished milestone.

● Upcoming milestone.

# AXSOME

## THERAPEUTICS

Thank you.

For more information, please contact

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