

**UNITED STATES
SECURITIES AND EXCHANGE COMMISSION**
Washington, D.C. 20549

Form 8-K

Current Report

Pursuant to Section 13 or 15(d) of The
Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): July 5, 2012

MEDICINOVA, INC.

(Exact name of registrant as specified in its charter)

DELAWARE
(State or other jurisdiction
of incorporation)

001-33185
(Commission
File Number)

33-0927979
(I.R.S. Employer
Identification No.)

**4350 LA JOLLA VILLAGE DRIVE,
SUITE 950, SAN DIEGO, CA**
(Address of principal executive offices)

92122
(Zip Code)

Registrant's telephone number, including area code: (858) 373-1500

Not applicable.

(Former name or former address, if changed since last report.)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions (see General Instruction A.2. below):

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Item 8.01 Other Events.

On July 5, 2012, MediciNova, Inc. (the "Company") updated the slide presentation to be used by the Company at investor meetings. A copy of the revised slide presentation is attached hereto as Exhibit 99.1 and is incorporated herein by reference.

Item 9.01 Financial Statements and Exhibits.

(d) *Exhibits.*

<u>Exhibit No.</u>	<u>Description</u>
99.1	Slide presentation of the Company.

SIGNATURE

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

MEDICINOVA, INC.

By: /s/ Michael Gennaro
Michael Gennaro
Chief Financial Officer

Date: July 5, 2012

EXHIBIT INDEX

Exhibit
No.

Description

99.1 Slide presentation of the Company.



*Accelerating
the global development
and commercialization of
innovative pharmaceuticals*



Forward-Looking Statements

Statements in this presentation that are not historical in nature constitute forward-looking statements within the meaning of the safe harbor provisions of the Private Securities Litigation Reform Act of 1995. These forward-looking statements include statements regarding MediciNova's clinical trials supporting the safety and efficacy of its product candidates and the potential novelty of such product candidates as treatments for disease, plans and objectives for clinical trials and product development, strategies, future performance, expectations, assumptions, financial condition, liquidity and capital resources. These forward-looking statements may be preceded by, followed by or otherwise include the words "believes," "expects," "anticipates," "intends," "estimates," "projects," "can," "could," "may," "will," "would," or similar expressions. Actual results or events may differ materially from those expressed or implied in any forward-looking statements due to various factors, including the risks and uncertainties inherent in clinical trials and product development and commercialization, such as the uncertainty in results of clinical trials for product candidates, the uncertainty of whether the results of clinical trials will be predictive of results in later stages of product development, the risk of delays or failure to obtain or maintain regulatory approval, the risk of failure of the third parties upon whom MediciNova relies to conduct its clinical trials and manufacture its product candidates to perform as expected, the risk of increased cost and delays due to delays in the commencement, enrollment, completion or analysis of clinical trials or significant issues regarding the adequacy of clinical trial designs or the execution of clinical trials and the timing, cost and design of future clinical trials and research activities; the timing of expected filings with the FDA; MediciNova's failure to execute strategic plans or strategies successfully; MediciNova's collaborations with third parties; MediciNova's ability to realize the anticipated strategic and financial benefits from its acquisition of Avigen, Inc., to integrate the two ibudilast development programs and to pursue discussions with potential partners to secure a strategic collaboration to advance the clinical development of the combined development program; the availability of funds to complete product development plans and MediciNova's ability to raise sufficient capital when needed, or at all; MediciNova's ability to comply with the covenants in its financing agreements; intellectual property or contract rights; and the other risks and uncertainties described in MediciNova's filings with the Securities and Exchange Commission, including MediciNova's annual report on Form 10-K for the year ended December 31, 2011 and its subsequent periodic reports on Forms 10-Q and 8-K. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of July 3, 2012. MediciNova disclaims any intent or obligation to revise or update these forward-looking statements.



Corporate Overview: MediciNova, Inc.

MediciNova Overview:

- Founded in September 2000
- Headquartered in San Diego, CA, with an office in Tokyo, Japan
- Dual listing on NasdaqGM as **MNOV** Osaka Securities Exchange as **4875**
- \$37.5 million market cap (NasdaqGM) as of 6/26/2012 (aggregate value of 18.3 million shares outstanding of common preferred on an as converted basis)

In-Licensed Clinical Stage Compounds:

- Unique access to differentiated, potentially high-value assets primarily from Japanese alliances (Kissei, Kyorin, Mitsubishi Tanabe Pharma, Meiji)

New Approaches to Treat Serious Medical Conditions:

- Bedoradrine Sulfate (**MN-221**): Intravenous treatment for acute exacerbations of asthma and chronic obstructive pulmonary disease (COPD)
- Ibudilast (**MN-166**): Oral treatment for progressive multiple sclerosis, neuropathic pain, and drug addiction



Business Model: Return On Investment

In-License:

- Novel, small-molecule product candidates with significant clinical or preclinical data packages and attractive market opportunities

Conduct Proof-of-Concept Clinical Trials:

- Conduct Phase 1 and Phase 2 clinical trials to demonstrate safety and efficacy of compound



Mitsubishi Tanabe Pharma

Two Pathways After Phase 2 (Proof-of-Concept):

- Internal development of compound towards commercialization in North America
- Seek partnership for further development of compound

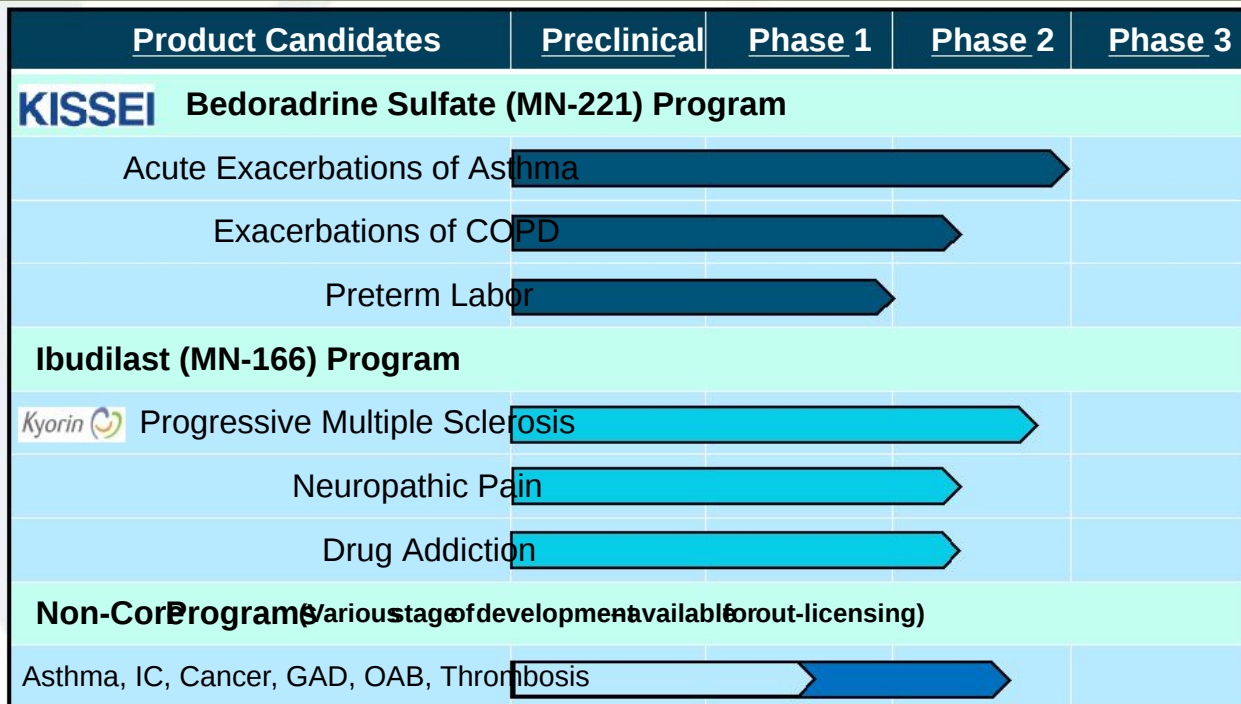


ANGIOGENE
PHARMACEUTICALS LTD

MEDICINOVA



Commercially-Attractive Diversified Portfolio





Significant Milestones

Milestone:

Timeline*:

- | | | |
|-------------------------------------|--|------------|
| <input checked="" type="checkbox"/> | Receive Use Patent for Ibudilast in Progressive MS Patients | 1Q, 2012 |
| <input checked="" type="checkbox"/> | Results from Phase 2b MN-221-CL-007 Acute Asthma Trial | 2Q, 2012 |
| <input type="checkbox"/> | Top-line Results from Phase 1b Multi-Dose Trial in COPD | 3Q, 2012 |
| <input type="checkbox"/> | Plan to Announce Phase 2 Clinical Program for Ibudilast (Asthma) | 3Q, 2012** |
| <input type="checkbox"/> | Plan to Announce Phase 2 Clinical Program for Ibudilast (MS) | 4Q, 2012** |
| <input type="checkbox"/> | End of Phase 2 Meeting with FDA for MN-221 Development | 4Q, 2012 |
| <input type="checkbox"/> | Commence Pivotal MN-221 Trial | 1H, 2013 |

**Anticipated completion dates based on current projections*

***Tentative based on availability of non-dilutive financing*

© MediciNova, Inc. 2012

MN-221:

- ***Acute Exacerbations of Asthma***
- ***Exacerbations of COPD***



Acute Exacerbations of Asthma (AEA)

Definition:

- Long-lasting and severe asthma episode that is not responsive to initial bronchodilator or corticosteroid therapy

Market Opportunity*:

- Approximately 1.9 million annual emergency room visits in the U.S.
 - ~500,000 annual hospitalizations in U.S. (~560,000 in UK/Spain/Germany/France/Italy)
 - Average length of stay for asthma hospitalization is 3.3 days (U.S.)
 - Average cost for asthma hospitalization is \$6,477
- Roughly 50% of subjects do not initially respond to standard care

Current Standard of Care (SOC):

- Inhaled beta agonists, inhaled anticholinergics, and IV or oral corticosteroids
- Current treatments are limited by **Bronchoconstriction** (insufficient airflow due to inflammation and airway constriction prevents inhaled drug uptake in the lungs) and **Mucus Plug Formation** (Persistent airflow limitation due to mucus secretion)

*Source: National Center for Health Statistics / CDC, WHO website, "Core Health indicators", 2007 National Hospital Discharge Survey, IMS Health's Disease and Condition Benchmarks – PharMetrics Integrated Database, 1/2007 – 12/2008
© MedicNova, Inc. 2012

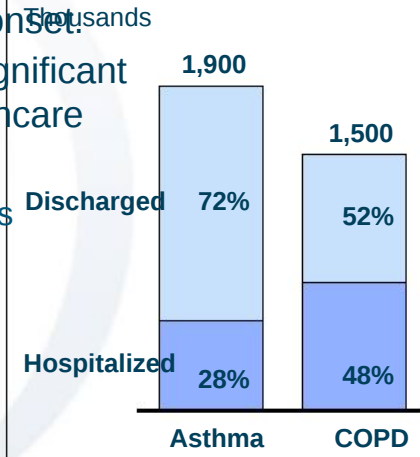


COPD Exacerbations

A COPD exacerbation is a sustained worsening of the patient's condition, from the stable state and beyond normal day-to-day variations, that is acute in onset. COPD exacerbations are associated with a significant increase in mortality, hospitalization and healthcare utilization.

- 1.5 million hospital emergency department visits
- 765,000 hospitalizations
 - Average length of stay 7.4 days*
 - Average cost ~\$32,000*
- 119,000 deaths

Hospitalization rates amongst Asthma and COPD patients



COPD patients are generally more ill than asthmatics with overall higher hospitalizations and mortality.

Source: CDC, CDC COPD surveillance in U.S.; US Census; American Lung Association website
*For COPD pts. with anemia; for pts. w/o anemia the stay is ~5.8 days at a cost of ~\$25K
© MediciNova, Inc. 2012





MN-221: A New Approach to Treating Acute Exacerbations (Asthma & COPD)

MN-221: A novel, highly selective β_2 -adrenergic receptor agonist

Potential advantages over current therapy:

1. Improved Efficacy

- Route of administration (IV vs. inhalation)
- Facilitates bronchodilation and onset clinical improvement

2. Improved Safety

- High selectivity for β_2 receptor versus β_1
- Partial agonist for β_1 receptor*

3. Reduced Health Care Expenses

- Reduction in hospitalizations; Return ER visits

** β_1 receptors are primarily responsible for cardiovascular stimulation*



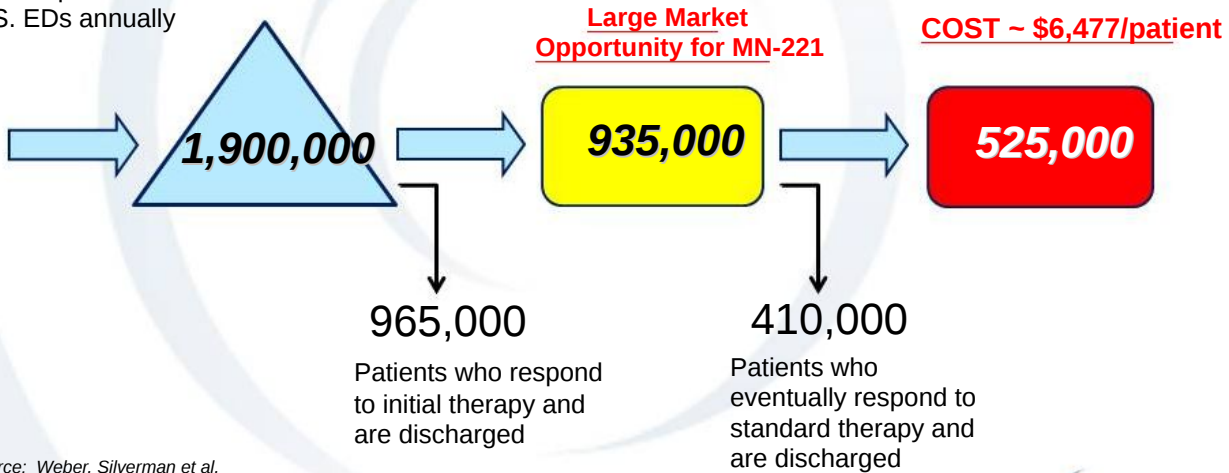
Acute Asthma Treatment Flow in Emergency Departments (EDs) in the U.S.

Input:
1,900,000 patients with acute exacerbations of asthma present at U.S. EDs annually

1st line therapy in ED:
Patients receive SOC, many while in the waiting room

2nd line therapy in ED:
Patients who do not initially respond continue receiving SOC

Hospitalization:
Patients who do not respond to SOC are eventually hospitalized



*Source: Weber, Silverman et al, American Journal of Medicine, Volume 113; pp 371

© MediciNova, Inc. 2012



MN-221 for Treatment of AEA: Pivotal Trial Development Strategy

End-of-Phase 2 Meeting with the FDA

- ✓ Scheduled for Monday, October 22, 2012

Pivotal Trial Design Modifications from Phase 2 Clinical Trial Based on these Core Principles*:

1. Primary Endpoints should be FEV₁ improvement at Hour 1 (delta) or AUC through Hour 2
2. Reduced variability (FEV₁ methodology, control for standard-of-care medications between study arms)
3. Larger sample size
4. Simpler Protocol for ease of enrollment
5. Include standardized clinical assessment at end of treatment period as secondary endpoint

*Tentative based on outcome of End-of-Phase 2 meeting with the FDA
© MediciNova, Inc. 2012



MN-221-CL-007: Phase 2 Trial Goals

Goals of the Phase 2b Clinical Trial:

1. Assess Efficacy adjunctive to SOC treatment
2. Establish Safety
3. Validate Proof-of-concept (POC) and Determine Pivotal Trial Design
4. Develop a basis for a successful End-of-Phase 2 Meeting with the FDA



MN-221-CL-007: Phase 2 Trial

Study Design

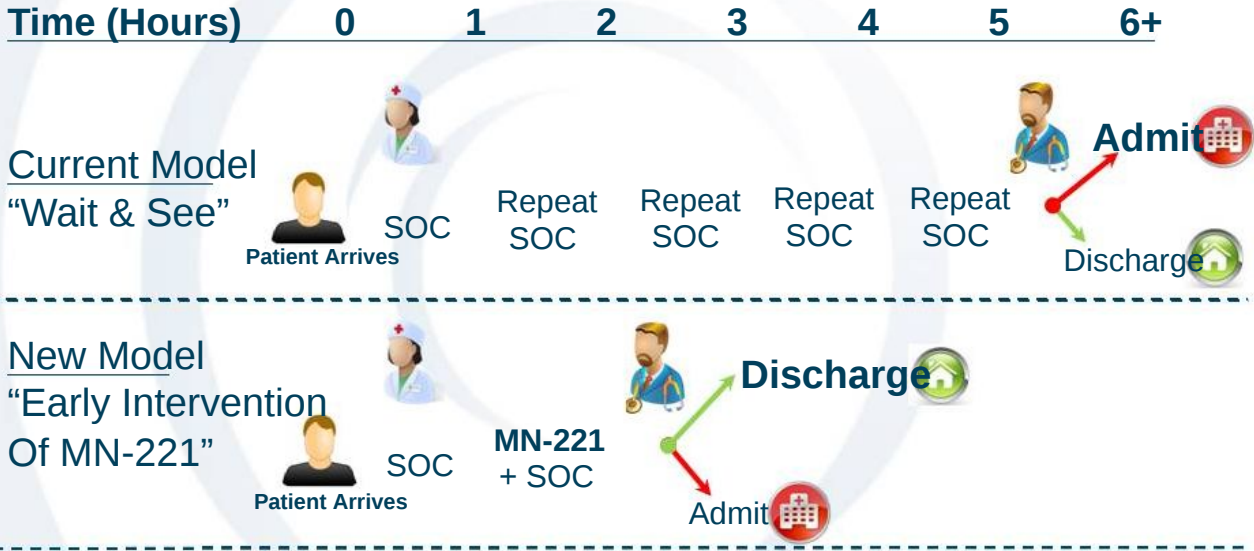
- Randomized, placebo-controlled, double-blind, multi-center Phase 2 clinical trial
- 175 patients enrolled with acute exacerbations of asthma at multiple US ED sites
 - 164 patients in the Efficacy Evaluable (EE) population; some patients early-terminated from the study
- Two treatment groups (1:1 randomization)
 - 1,200µg infusion of MN-221 over 1hr + Standard-of-Care (SOC)
 - Placebo infusion + Standard-of-Care
- Primary Efficacy Endpoint was AUC of change in FEV₁ hours 0 - 3
- Important Secondary Endpoints include:
 - Improvements in FEV₁ at other time points
 - Clinical improvement outcomes: Dyspnea score, Respiratory Rate
 - Pharmacoeconomic benefits: Hospitalization admissions and Return ER visits*

*As captured in the seven-day patient follow-up; not official secondary endpoint per protocol.



MN-221:

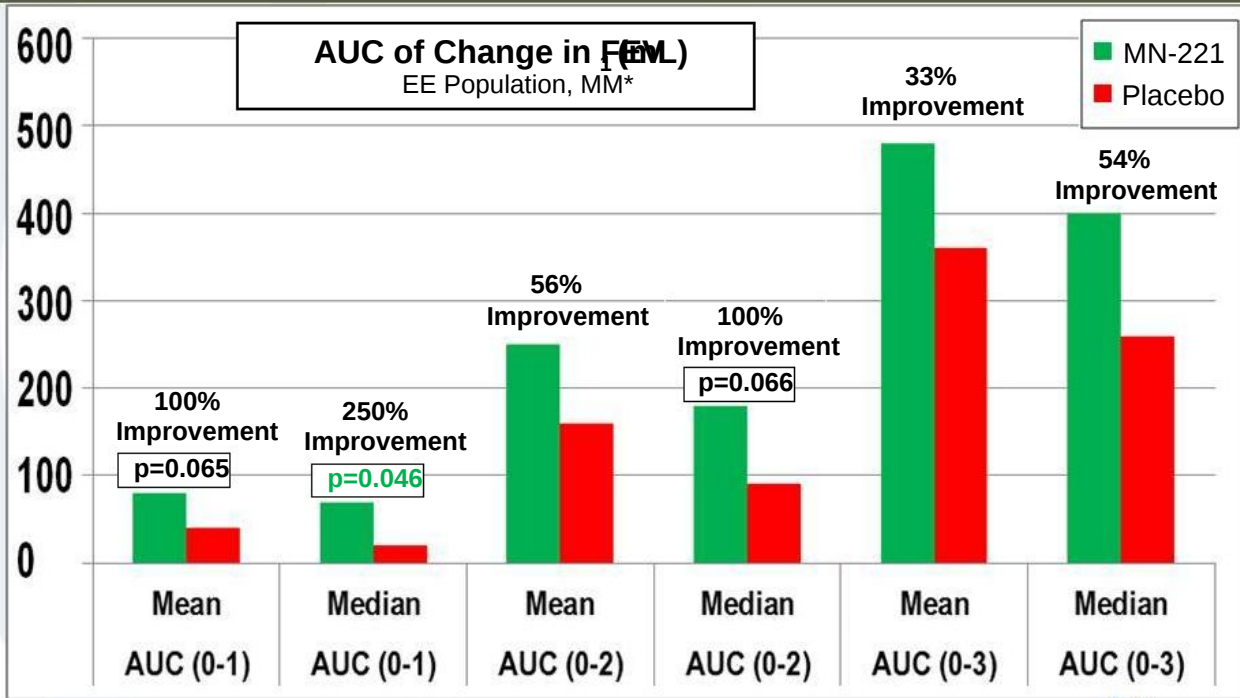
Potential New Model for Treating AEA in the ER



- ❖ ER cost per hour per patient is expensive
- ❖ The longer the patient is in the ER the greater the probability of admission



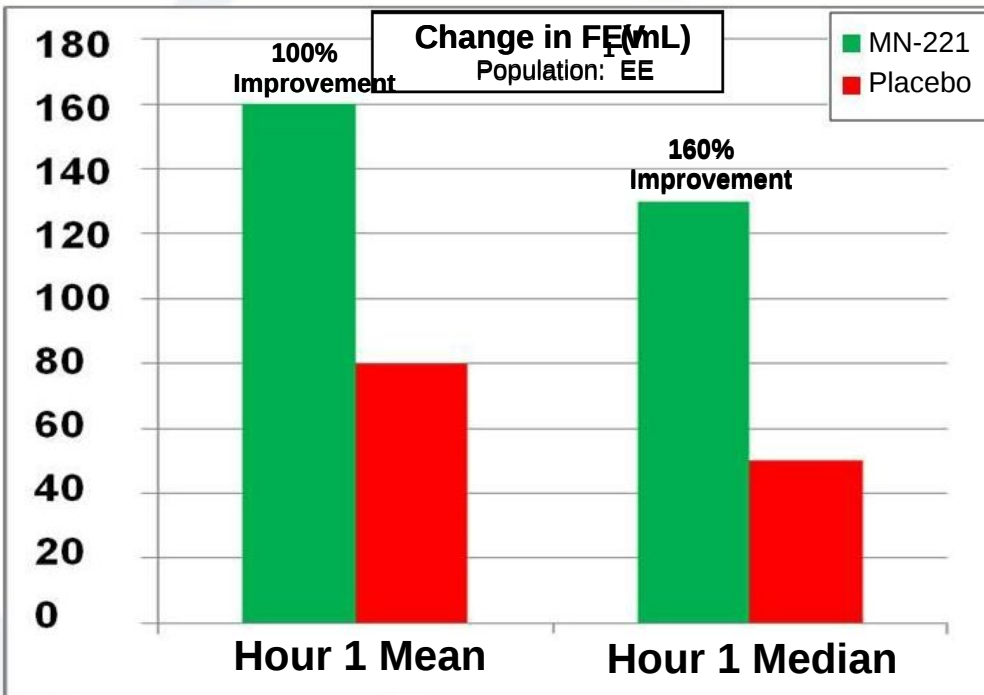
MN-221-CL-007: Efficacy Endpoints for FEV₁



*EE = Efficacy Evaluable Population; MM= Mixed Model was used for statistical analysis of efficacy parameters
© MediciNova, Inc. 2012

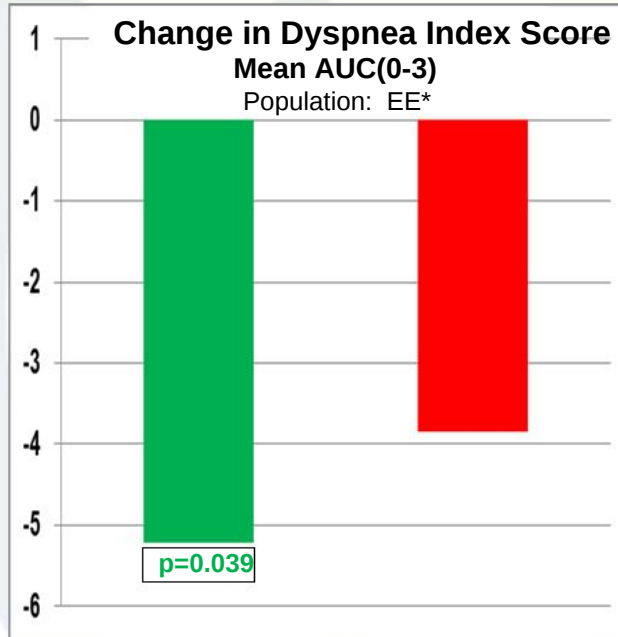


MN-221-CL-007: Change in FEV₁ at Hour 1



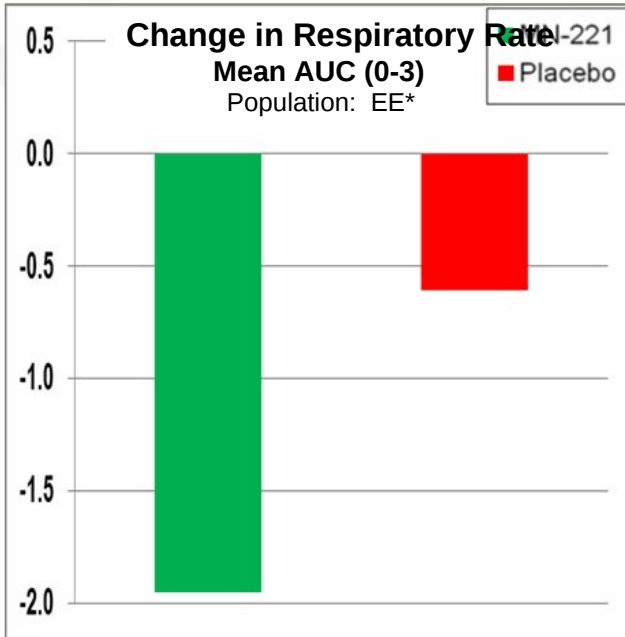


MN-221-CL-007: Improvements in Clinical Symptoms



Units are dyspnea index score
(0-10 scale; 5=severe, 3=moderate, 2=slight)

© MediciNova, Inc. 2012



Units are breaths/min.

*EE = Efficacy Evaluable Population





MN-221-CL-007: Analysis of High Responders

Analysis of High Responders:

FEV₁ Improvement at Any Time Point during the Treatment Period
EE population*

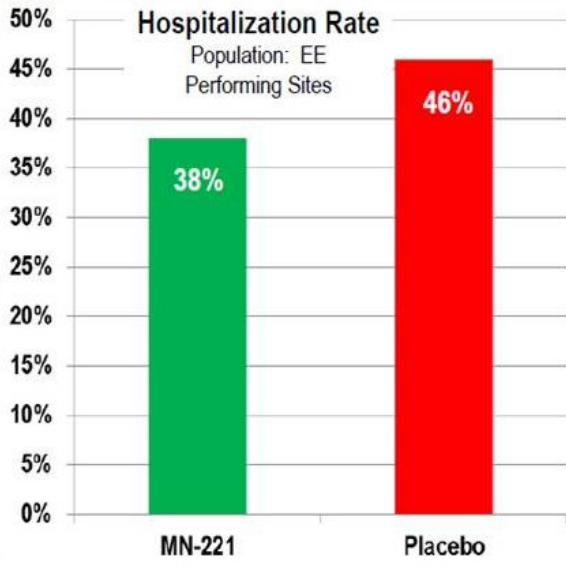
FEV ₁ Improvement	Placebo (%)	MN-221 (%)	P-value
≥ 200mL	37 / 77 (48%)	48 / 78 (62%)	0.09
≥ 10% pred.	25 / 77 (32%)	34 / 78 (44%)	0.15

*EE = Efficacy Evaluable Population

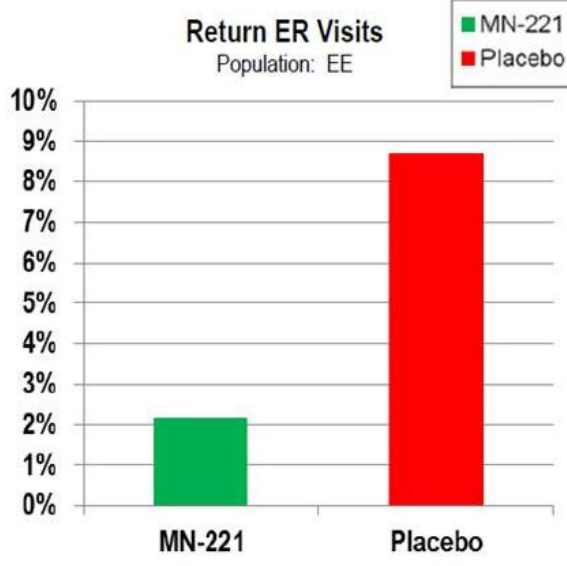


MN-221-CL-007: Phase 2 Trial

Pharmacoeconomic Benefits



The addition of MN-221 resulted in a 17% reduction in hospital admissions*



A higher percentage of patients in the placebo group returned to the ER within 7 days

*EE = Efficacy Evaluable Population; Performing Sites analysis includes patients from sites completing more than 2 patients during the trial and for which all efficacy measurements were available.



MN-221-CL-007: Phase 2 Trial

MN-221 Met its Goals of the Phase 2 Program

1. Assess Efficacy adjunctive to SOC treatment

- ✓ MN-221 group showed improvement over placebo group in measurements of lung function
- ✓ MN-221 showed a notable reduction in hospital admissions in performing sites and a notable reduction in ER return visits

2. Establish Safety

- ✓ No Serious Adverse Events related to MN-221; >400 subjects exposed

3. Affirm POC and guides Trial Design for Pivotal Trials

- ✓ Endpoint modifications, Larger 'n', reduced variability with protocol and standard-of-care treatments, and clinical assessment of improvement at end of treatment

4. Develop a basis for a successful End-of-Phase 2 Meeting with the FDA

- ✓ Meeting Scheduled for October 22, 2012



MN-221-CL-012:

Ongoing Phase 1b/2a COPD Trial

- Randomized, placebo-controlled, double-blind, Phase 1b/2a clinical trial
 - ~20 stable moderate-to-severe COPD patients (FEV₁ 30-80%)
 - Two treatment groups
 - 1,200µg infusion of MN-221 every 12/24 hours (15 pts.) over 4 days
 - Placebo (5 pts.)
 - Primary objective is to determine the safety and tolerability of administered **multiple times** over several days in COPD patients who may also have **co-morbidities and concomitant medications** common in this population.
 - Secondary objectives include pharmacokinetics, preliminary efficacy of repeated administration of MN-221 in COPD patients, and testing of a simple, hand-held digital FEV₁ & Peak Flow monitoring device.
- **Top-line results expected 3Q, 2012**

MN-221:

- ***Market Opportunity***
- ***Patent Summary***
- ***Next Steps***



MN-221 Market Opportunity*

Market	Acute Asthma	COPD Exacerbations
US	\$375-400 million	\$380-420 million
Europe	\$200-300 million	\$200-300 million
Rest of World	\$150-250 million	\$150-250 million
Worldwide MN-221 Sales Potential	\$725-950 million	\$730-970 million
Combined Worldwide MN-221 Sales Potential	\$1.5-1.9 Billion	

**Prices in today's dollars, do not reflect any price increases which may be implemented;
Assumes a conservative price per dose target ~\$550/dose*

Source: Physician interviews, team analysis



MN-221 Patent Summary

- The U.S. patent for MN-221 has composition of matter and method of use claims and is set to expire no earlier than February 2017
 - Corresponding composition of matter patents in various other countries
 - U.S. patent expiration does not include Waxman-Hatch patent term restoration (industry average = 4.5 years)
 - Waxman-Hatch grants 5 years of exclusivity from approval in the U.S. (*We anticipate this along with pediatric exclusivity and ANDA review time will give us at least 7.0 years of exclusivity*)
- Exclusivity in Europe is 10 years for first approval of new chemical entities
- In addition, MediciNova has filed multiple patent applications related to MN-221 which if granted, could protect MN-221 until at least 2030



MN-221 Next Steps

1. Complete COPD Phase 1b/2a Clinical Trial
 - Important Safety data of multiple infusions
 - Validation of hand-held spirometry device
2. Market Analysis
 - Collaborate with leading market research firm to quantify the value of reduced hospital admissions
 - Quantify the value of reduced ER visits
3. End-of-Phase 2 Meeting with the FDA
 - Scheduled for Monday, October 22nd, 2012
 - Confirm endpoints and trial design for pivotal program; Review overall development plan
4. Strategic Partnership Discussions Ongoing

Ibudilast (MN-166/AV-411):

- ***Progressive Multiple Sclerosis***
- ***Neuropathic Pain***
- ***Addiction***



Ibudilast (MN-166) Overview

Ibudilast (MN-166)

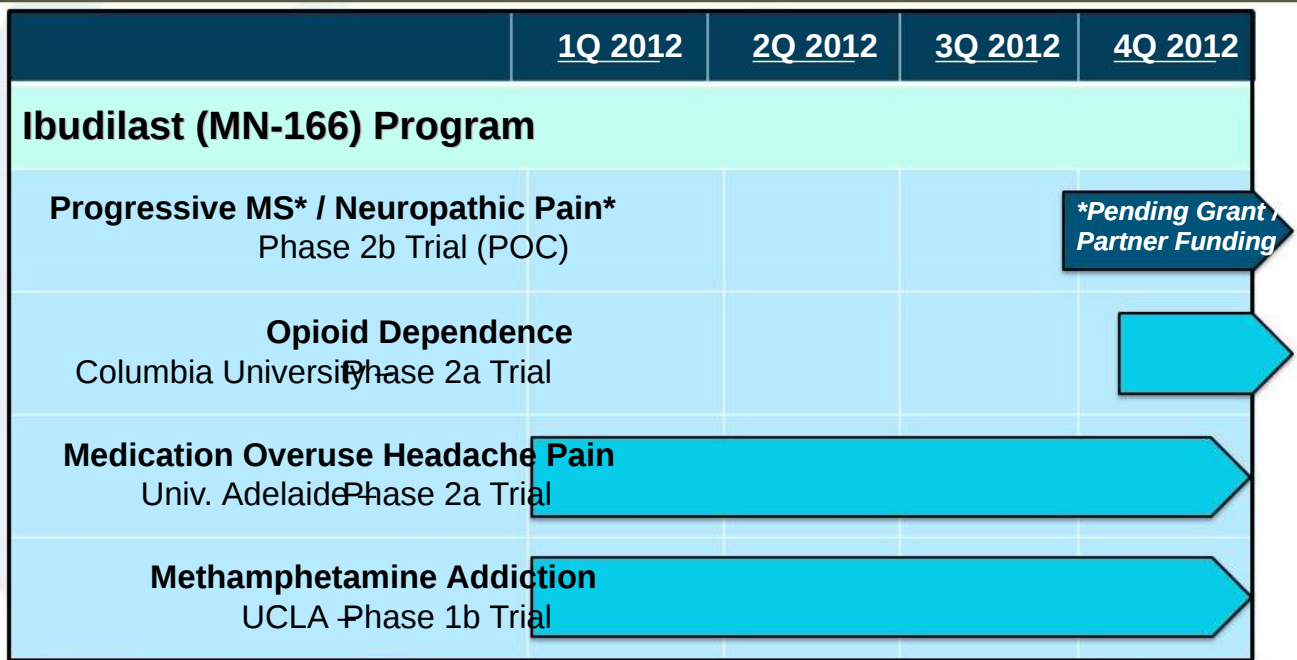
- Oral administration
- Safe and well tolerated
 - Approved in Japan/Korea over 3.2M patient exposures
 - >420 subjects treated with ibudilast
 - Dosing up to 100 mg daily doses
- Mechanism(s) of action primarily:
 - Inhibition of macrophage Migration Inhibitor Factor (MIF)
 - PDE-4,10 inhibition
 - Neurotrophic action and attenuation of glial cell activation

Clinical Safety & Preliminary Efficacy Established

- Phase 2 multiple sclerosis proof-of-concept study
 - Indicators of dose-related neuroprotective efficacy validated
- Phase 1 dosing to 100 mg/d
- Phase 1b/2a trial in diabetic neuropathic pain completed
- Phase 1b/2a clinical trial in opioid withdrawal & analgesia completed



Ibutilast (MN-166) Program: Ongoing and Future Development



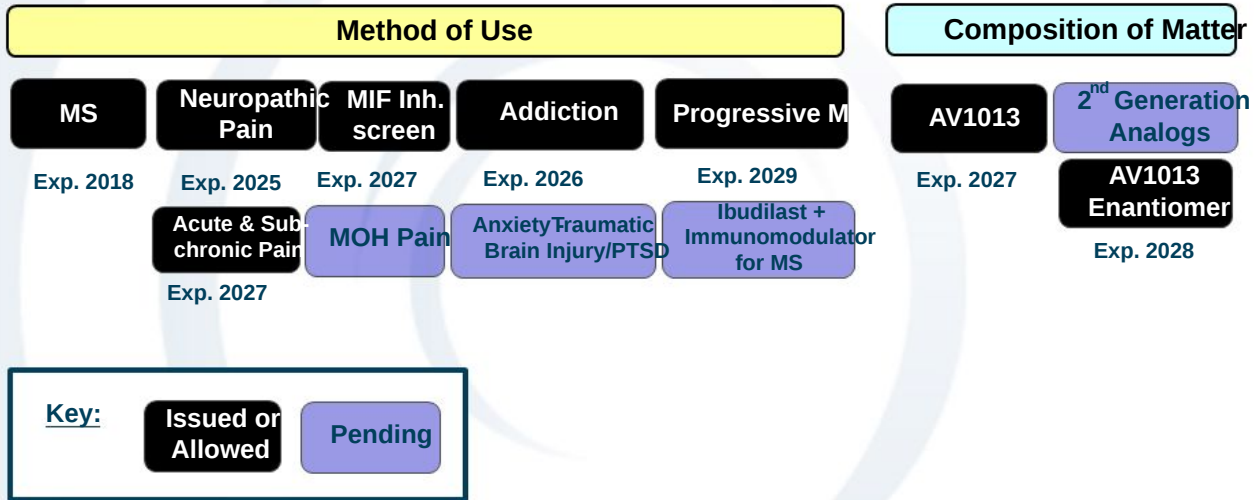


Strategy for Ibudilast's Development

- **Sustain NIDA-sponsored Drug Addiction Development**
 - Potential for Phase 2 POC Trial support
- **Potential for Gov't and MS Society consortium funding of Phase 2 Progressive MS POC Trial**
- **Collaboration with Development Partner:**
 1. Advance to Phase 2b Proof-of-Concept in MS and/or Pain
 2. Provide competitive economics for first in class therapy
 3. Could be collaboration through:
 - i. Pharma partner
 - ii. Project financing
 - iii. Shared-risk with competitive CRO agreement
- **Consider Investigator-sponsored Neurological Trials in Focus Areas**



Patent/Commercial Overview





Management Team with Global Experience



Leadership	Years Experience	Background
Yuichi Iwaki, MD, PhD CEO & President	35	Professor at USC, formerly Professor at University of Pittsburgh; Advisor to JAFCO, Tanabe
Michael Coffee Chief Business Officer	26	Avigen, Amarin Corp., Elan Pharmaceuticals, N.A., Athena Neurosciences
Kirk Johnson, Ph.D. Chief Scientific Officer	21	Avigen, Genesoft Pharmaceuticals, Chiron Corporation (Novartis San Francisco)
Masatsune Okajima, CMA VP, Head of Japanese Office	19	Daiwa Securities SMBC, Sumitomo Capital Securities, Sumitomo Bank
Kazuko Matsuda, M.D., Ph.D., MPH Chief Medical Officer	20	Assistant Professor USC, Keck School of Medicine; Children's Hospital Los Angeles.
Michael Gennaro, CPA, MBA Chief Financial Officer	37	Partner at FLG Partners, Sylanro Systems, Inverse Network Technology, Novell, Piiceon, Verticom



Financial Overview

Financial Resources:

- **\$11.0 million** in cash & cash equivalents as of 3/31/2012
- Including \$10 million raised through equity sale and non-dilutive funding by Kissei
 - \$7.5 million raised in private stock sale to Kissei Pharmaceutical Co., Ltd.
 - \$2.5 million additional non-dilutive R&D funding from Kissei
- ~18.3 million shares outstanding on an as converted basis
- Cash Runway into 2013



MediciNova Corporate Summary

Experienced Management Team

- ❖ Translational Medicine & Clinical development expertise
- ❖ Strong international presence, especially Japan
- ❖ Large and small pharma/biotech experience



Capitalization

- Raised ~\$18M in 2011
- Cash Runway into Q2, 2013

2012 Milestones

1. Announced Results from CL-007 Q1 -
2. Announce Results from CL-012 Q3 -
3. Announce Phase 2 Trial(s) for Ibudostat -
4. End-of-Phase 2 Meeting with FDA -

**Anticipated completion dates based on current projections*

© MediciNova, Inc. 2012

Addendum

Data from Completed Trials

➤ ***MN-221***

➤ ***Ibudilast (MN-166/AV411)***

MN-221:

Data from Completed Trials

- ***Asthma Program:***
 - ***CL-004, CL-005, CL-006, CL-007***
- ***COPD Program:***
 - ***CL-010***
- ***Safety Review***



MN-221 Clinical Development

Acute Asthma Program:

- Multiple doses tested at infusion lengths of 15min, 1hr, and 2hr
- Completed 2 trials in asthmatics with stable disease
 - **CL-004**(23patients)
 - **CL-005**(17patients)
- Completed Phase 2a trial in patients with AEA in the ED
 - **CL-006**(29patients)
- Completed Phase 2b study in patients with AEA in ED
 - **CL-007**(175pts.)
- End of Phase 2 Meeting with FDA
 - Scheduled for **October 22nd**

COPD Program:

- Multiple doses tested at 1 hour infusion
- Completed 1 trial in COPD patients with stable disease
 - **CL-010**(48patients)
- Preparing to initiate multi-dose trial in patients with stable COPD
 - **CL-012**(20patients)
 - Efficacy and Safety data will also be very useful in further development of MN-221 for acute asthma

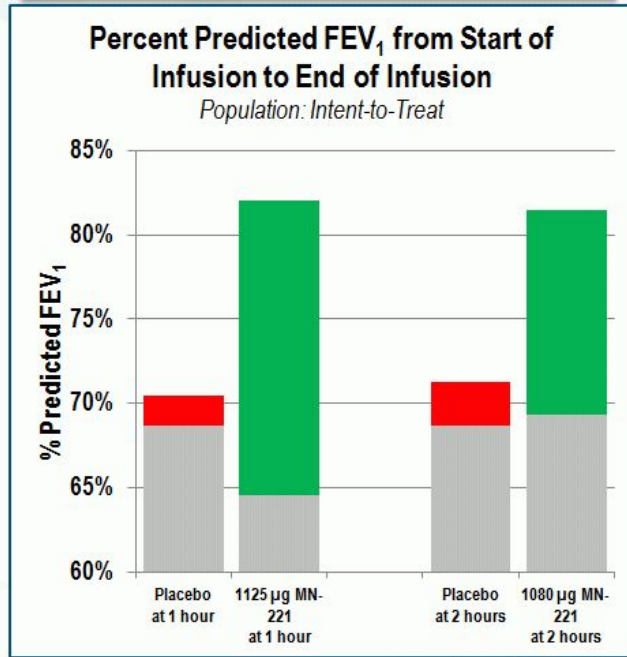
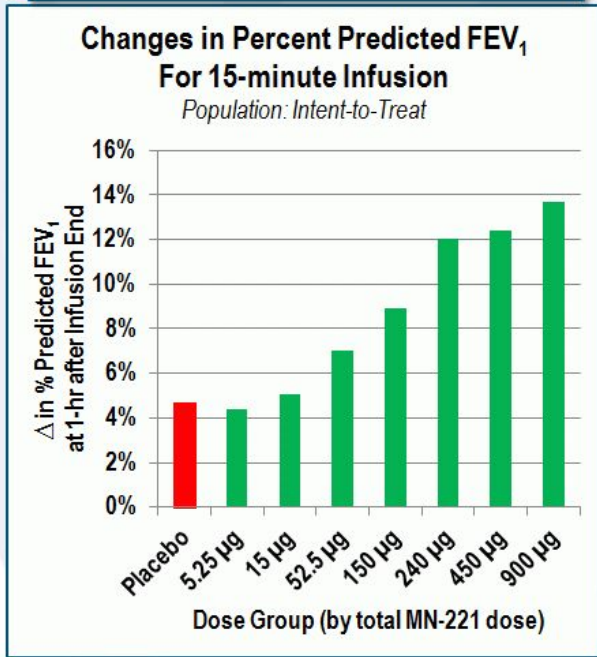


MN-221 Clinical Results

Improved Lung Function at Different Dosing Levels: Stable Asthmatics

CL-004: Stable Mild/Moderate Asthmatics

CL-005: Stable Moderate/Severe Asthmatics

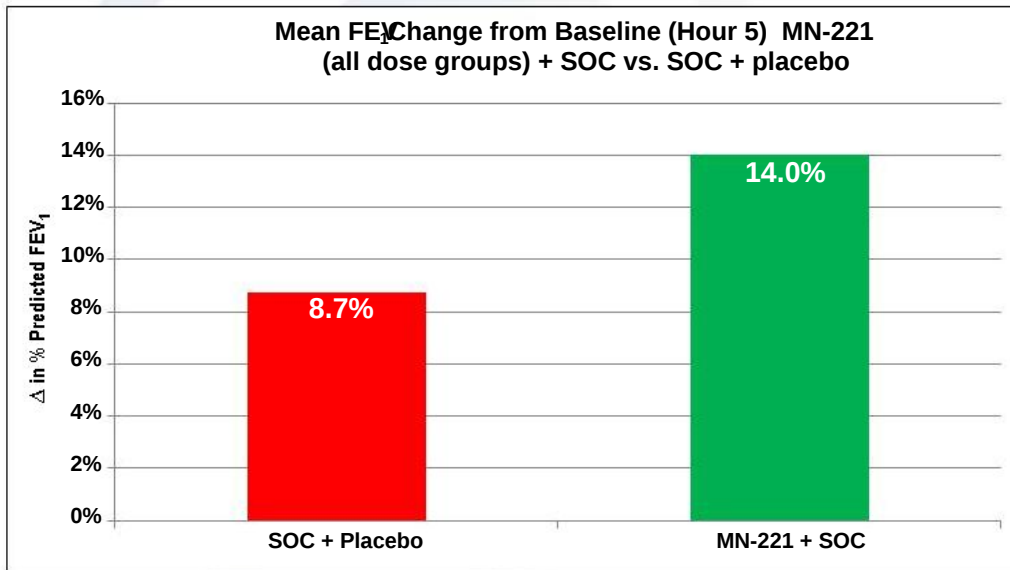




MN-221 Clinical Results

Improved Lung Function and Clinical Outcome Above and Beyond Standard of Care (SOC)

CL-006: Patients Suffering from Acute Exacerbation of Asthma in Emergency Department



Mean change in FEV₁ from baseline was 5.3% higher in the MN-221 dose groups versus the placebo group



MN-221-CL-007: Phase 2 Trial

Subject Populations

	Placebo	MN-221
Enrolled subjects	86	89
Safety population	84	83
EE population	83	81
Performing Sites population	70	72

Definitions of Note:

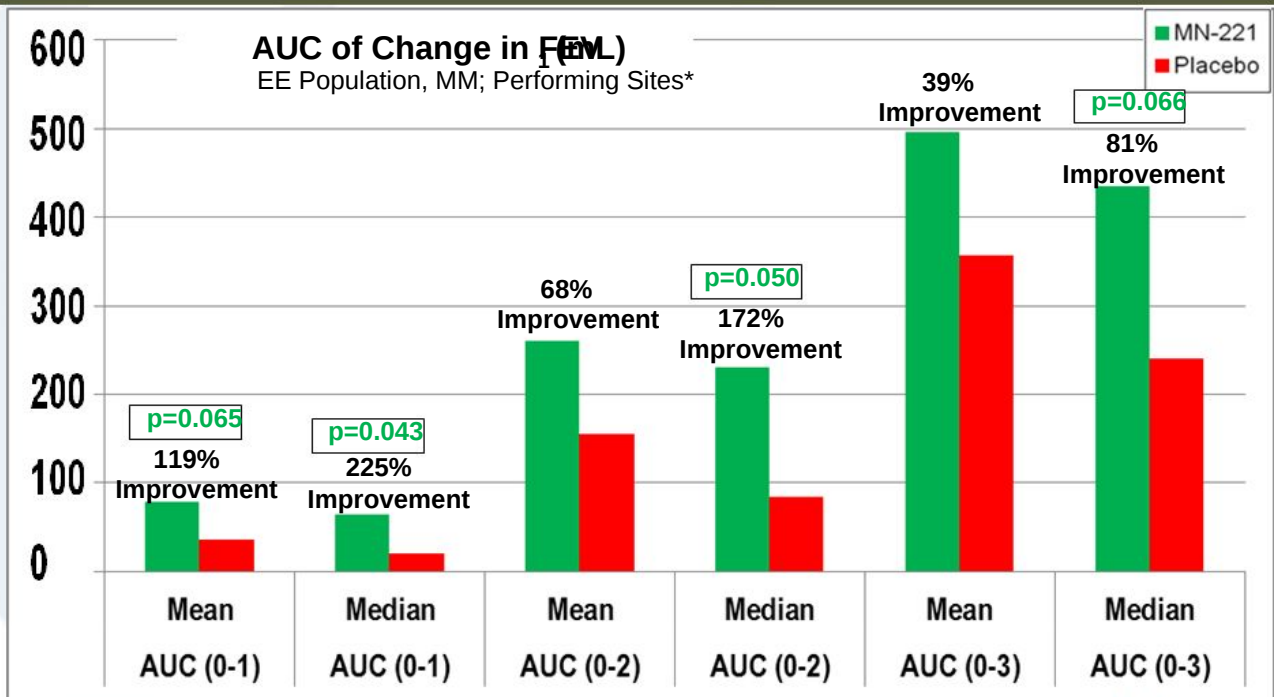
EE = Efficacy Evaluable Population

Performing Sites Population: analysis includes patients from sites completing more than 2 patients during the trial and for which all efficacy measurements were available.



MN-221-CL-007: Phase 2 Trial

Efficacy Endpoints for FEV₁

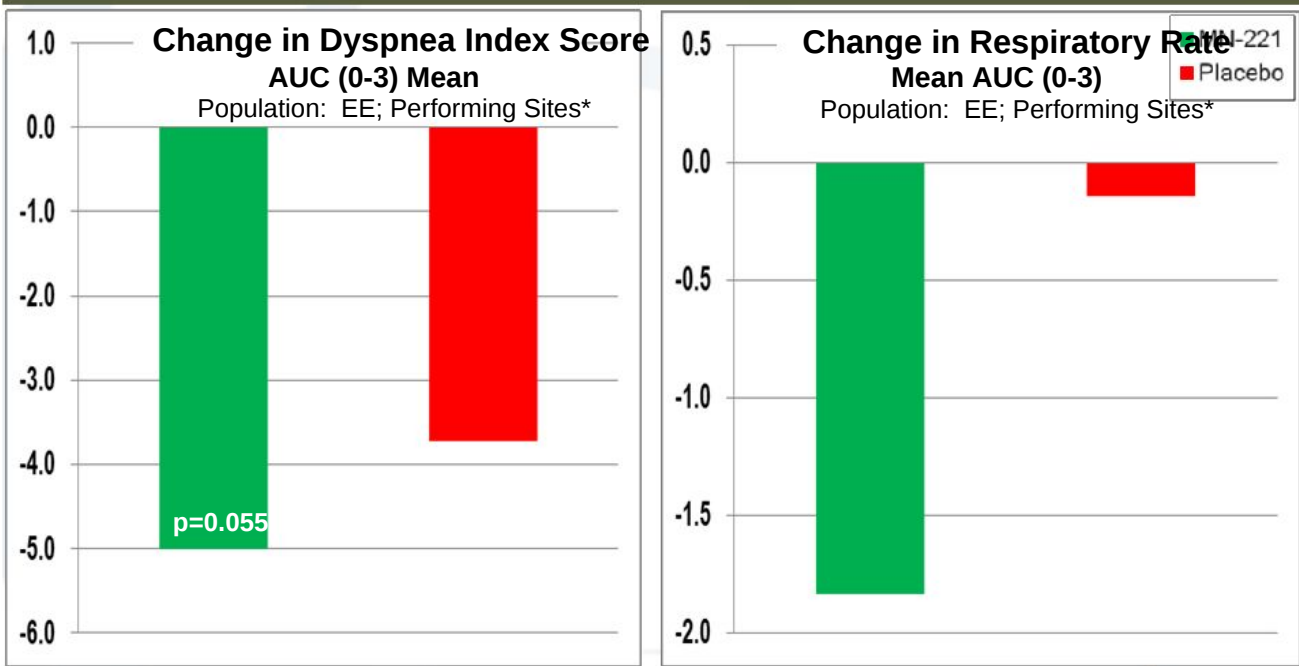


*EE = Efficacy Evaluable Population; MM= Mixed Model was used for statistical analysis of efficacy parameters; Performing Sites analysis includes patients from sites completing more than 2 patients during the trial and for which all efficacy measurements were available.



MN-221-CL-007: Phase 2 Trial

Improvements in Clinical Symptoms



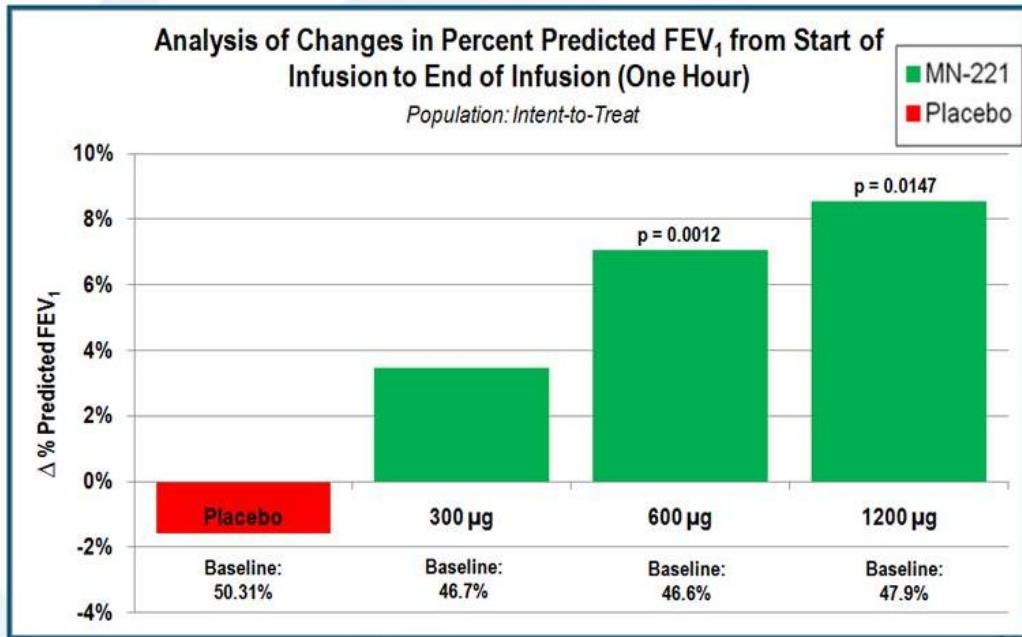
*EE = Efficacy Evaluable Population; Performing Sites analysis includes patients from sites completing more than 2 patients during the trial and for which all efficacy measurements were available.



MN-221 Clinical Results

Improved Lung Function at Different Dosing Levels: COPD Patients

CL-010: Stable Moderate/Severe COPD Patients





MN-221 Safety Review

- MediciNova has preclinical data and clinical data which demonstrates the safety of MN-221.
- In summary, we have not seen clinically significant safety concerns with MN-221 and it has been tested in over 400 subjects to date.
- According to interviews of emergency room physicians, less-selective injectable betaagonists such as epinephrine and terbutaline are not commonly used to treat acute asthma. The main reason they are not used more often is due to safety concerns, particularly cardiovascular side effects.



Human β -Adrenergic Receptor Selectivity

Test Drug	β_1 IC ₅₀ (M)	β_2 IC ₅₀ (M)	β_2 -Adrenoceptor Selectivity (IC ₅₀ for β_1 / IC ₅₀ for β_2)
Levalbuterol	7.40E-06	1.40E-06	5.3
Albuterol	9.40E-06	1.60E-06	5.9
Terbutaline	6.00E-05	6.50E-06	9.2
MN-221	5.90E-06	1.40E-07	42.4



MN-221: CHEST Posters (Nov. 2010)*

- **MN-221-CL-004** Evaluation of MN-221 (bedoradrine), Novel Highly Selective Beta2-Adrenergic Receptor Agonist in Mild to Moderate Asthma via Intravenous Infusion (Poster #145)
- **MN-221-CL-005** Comparison of Administration Rates of MN-221 (bedoradrine), Novel Highly Selective Beta2 Receptor Agonist in Patients with Stable Moderate to Severe Asthma (Poster #143)
- **MN-221-CL-006** Reduce Hospital Admission and Improve Pulmonary Function Following Intravenous MN-221 (bedoradrine), a Novel Highly Selective Beta2-Adrenergic Receptor Agonist, Adjunctive to Standard of Care in Severe Acute Exacerbation of Asthma (Poster #144)
- **Pharmacokinetic (PK) and Pharmacodynamic (PD) Modeling and Simulation Support the Novelty of MN-221, a Highly-Selective Beta2-Adrenergic Agonist for Treatment of Acute Asthma (Poster #146)**
- **MN-221-Y08-065** Cardiovascular Effects of i.v. MN-221 (bedoradrine) Administered with Nebulized Albuterol in Dogs (Poster #147)
- **Pharmacokinetics and Pharmacodynamics of MN-221, a Novel Highly-Selective Beta2-Adrenergic Agonist for Treatment of Acute Chronic Obstructive Pulmonary Disease (Poster #685)**
- **MN-221-CL-010** Intravenous MN-221, a Novel Highly Selective Beta2 Adrenergic Receptor Agonist, Improves Lung Function in Stable Moderate to Severe Chronic Obstructive Pulmonary Disease (COPD) Patients (Poster #686)

*Posters available on MediciNova website at www.medicinova.com

Ibudilast (MN-166):

Data from Completed Trials

- ***Multiple Sclerosis Program***
- ***Neuropathic Pain Program***
- ***Drug Abuse/Addiction Program***
- ***Safety Review***



Multiple Sclerosis Clinical Study: MN-166-CL-001

Placebo-controlled, Randomized, Double-blind Phase II Study:

- Year 1 Placebo, 30 mg/day, 60mg/day
- Year 2 30 mg/day, 60mg/day
- 297 patients (~100 patients/group) @ 25 sites in Serbia, Ukraine, Belarus, Bulgaria and Romania

Key Inclusion Criteria:

- Males or females aged 18 to 55 years, with relapsing remitting (RR) and/or secondary progressive (SP) Multiple Sclerosis with continued relapses;
- One MRI scan taken two weeks prior to treatment start using a standardized MRI protocol with at least one Gd-enhancing lesion;
- An Expanded Disability Status Scale (EDSS) score of 5.5 or less at the screening and baseline visits.

Safety Profile:

- 89% (264 of 297) of subjects completed the first 12 months of the study
- 82.5% (245 of 297) of subjects completed the full 24 months of the study
- Adverse effects reported more frequently in MN-166-treated than placebo-treated subjects: GI effects & depression



MN-166-CL-001 Study Results

Indicative of Potential Neuroprotective Effect:

- Reduced brain volume loss **P-Value: 0.035**
- Reduced conversion of acute lesions to persistent black holes **P-Value: 0.004**
- Sustained disability progression was significantly less likely ($\approx 50\%$) **P-Value: 0.026**

Acute Clinical Benefit:

- Prolong time to relapse (by 127 days.) **P-Value: 0.04**
- Annualized relapse rate **P-Value: 0.08**

Protocol-Defined Primary Endpoint (Surrogate Endpoint):

- No significant reduction in the cumulative number of active (gadolinium-enhancing (T1) and non-enhancing new/enlarging (T2)) lesions on cranial MRI scans over 12 months of treatment was observed
- Positive trends were observed in volume of gadolinium-enhancing (T1) lesions **P-Value: 0.09**

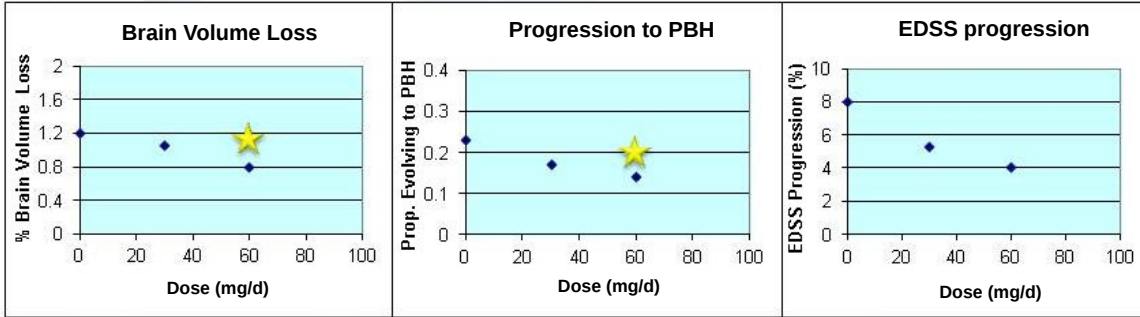
Note: P-values listed on this slide compare placebo group to 60mg/day group of MN-166

© MediciNova, Inc. 2012

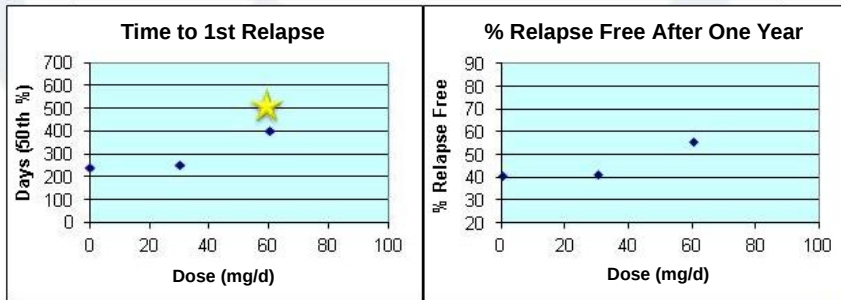


MN-166-CL-001: Efficacy Review (One Year)

Endpoints Indicative of Disease Modifying Effect (Chronic aspects of MS):



Endpoints Relating to Acute Clinical Benefit:



★ : Statistically significant



Secondary Progressive MS: Subset Analysis

Subset of MS Patients:	Treatment Group (patient n)				
	Placebo	30 mg/day Low Dose Group		60 mg/day High Dose Group	
	% Brain Volume Change	% Brain Volume Change	Magnitude of Effect	% Brain Volume Change	Magnitude of Effect
RRMS	-1.2 (81)	-1.1 (69)	8% less	-0.8 (75)	33% less
SPMS	-1.0 (3)	-0.7 (4)	30% less	-0.4 (2)	60% less

Next Steps for Progressive MS:

Two-year Phase 2 in Progressive MS - month 12 data. Potential first-in-class once or twice-daily oral well-tolerated drug with established endpoints. Draft protocols, costs and trial operations completed.



Diabetic Peripheral Neuropathic Pain Study: AV411-010

Design: Two-center (Australian), Phase 1b/2a, randomized, double-blind, placebo-controlled, parallel-group study.

Subjects:

- Patients, aged 18 to 75 years, with painful diabetic peripheral neuropathy (DPN) or complex regional pain syndrome (CRPS) of ≥ 6 months duration and screening VAS score ≥ 4 cm on a 10 cm scale
- 29 subjects: 19 active, 10 placebo

Dosing:

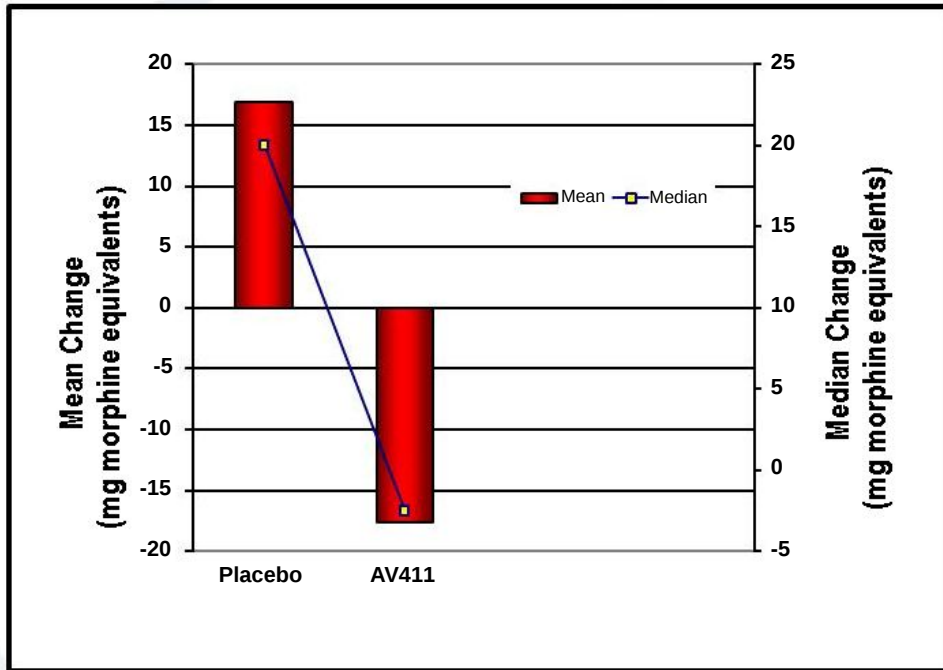
- 20 mg BID (n=4), 20 mg TID (n=4), 40 mg BID (n=11)
- AV411 (ibudilast) added to patients' standard medication regimen for DM and *pain*

Study objectives:

- Establish safety/tolerability & PK in intended patient population
- Explore potential efficacy endpoints



Reduction Observed in Opioid Usage





Greater % of “Responders” Above Ibudilast Plasma Thresholds

	Plasma Ibudilast Parameter	VAS ‘Responder’ %
AUG_{0-24h}	> 1000 ng*hr/mL	60%
	< 1000 ng*hr/mL	25%
C_{max}	> 60 ng/mL	64%
	< 60 ng/mL	14%
C_{min}	> 27 ng/mL	55%
	< 27 ng/mL	29%

Next Steps for Neuropathic Pain:

Twelve week Phase 2 DPN trial. Potential first-in-class, once- or twice-daily oral well-tolerated drug with established endpoints. Draft protocols, costs and trial operations completed.



Investigator-Led Development

- Recent validation of CNS action in opioid withdrawal & analgesia
- Ongoing Methamphetamine interaction Phase 1b
- Opioid self-administration Phase 2a initiating
- Ongoing Phase 2a Medication Overuse Headache Pain trial
 - Randomized, double-blind, placebo-controlled, investigator-initiated (Dr. Pail Rolan at Univ. of Adelaide, Australia; reduced headache index, acute medication (codeine) use and headache impact on Quality of Life (QOL); 8-week trial + follow-up; n = 20 patients each at placebo vs. 80 mg/day of MN-166
- Acquired rights to treatment of post-traumatic brain injury (TBI)
 - Led by the research of Daniel Barth, Ph.D., Professor of Neuroscience and Psychology at CU-Boulder, ibudilast demonstrated significant efficacy in a model of post-TBI anxiety, one of the most common disorders caused by TBI.



Ibutilast (MN-166): Neurological Indications and Translational Record

Indication	Preclinical Validation	Clinical Validation	Comment
MS	+ (EAE)	+* MN-166-CI-001	Progressive MS Phase 2b indicated
Neuropathic Pain	+ (multiple models)	+ AV411-010	Phase 2b enabled
Opioid Dependence (and Tolerance)	+ (multiple models)	+* AV411-OWA	(SOWS, Miosis)
Enhanced Opioid Analgesia	+ (2 rat models)	+* AV411-OWA	(McGill PQ)
Methamphetamine Relapse	+ (rat models)	in progress	
Traumatic Brain Injury+ (rat models)		TBD	

* = $p < 0.05$, dose-related & certain endpoints



Ibudilast - References

- Barkhof, F. et al. *Ibudilast in Relapsing-Remitting Multiple Sclerosis: a Neuroprotectant?* *Neurology*, Mar 30 2010.
- Fox, R. *Primary Neuroprotection: the Holy Grail of Multiple Sclerosis Therapy.* *Neurology*, Mar 30 2010.
- Kagitani-Shimono K. and Mohri I. *J Neuroinflammation. Anti-inflammatory Therapy by Ibudilast, a Phosphodiesterase Inhibitor, in Demyelination of Twitcher, a Genetic Demyelination Model of Inflammation.* 2005; 2(1): 10.
- Kreutzberg G. W. *Microglia: A Sensor for Pathological Events in the CNS.* *Trends Neurosci.* 1996; 19(8): 312-8.
- Ledeboer A., Hutchinson M. R., Watkins L. R., and Johnson K. W. *Ibudilast (AV411): A New Class Therapeutic Candidate for Neuropathic Pain and Opioid Withdrawal Syndromes.* *Investigative Drugs* 2007; 16:935-950.
- Mizuno, T et al. *Neuroprotective role of phosphodiesterase inhibitor ibudilast on neuronal cell death induced by activated microglia.* *Neuropharmacology* 46:404, 2004.
- Muzio L., Martino G., et al. (2007). *Multifaceted Aspects of Inflammation in Multiple Sclerosis: The Role of Microglia.* *J Neuroimmunol* 2007; 191(1-2): 39-44.
- Rolan, P., Hutchinson, M., and Johnson, K. *Ibudilast: A Review of its Safety, Efficacy, and Pharmacology in Respiratory and Neurologic Diseases.* *Expert Opinion Pharmacotherapy* 2009.
- Wang, F. et al. *Spinal Macrophage Migration Inhibitory Factor Is a Major Contributor to Rodent Neuropathic Pain-like Hypersensitivity.* *Anesthesiology* 2011Feb2.