

Final Outcomes Summary: Live Webinar and Twitter Chat Journal Club

**August 2021 – August 2022** 

**Grant ID: 67929297** 

Supported by an Educational Grant from Insmed.



**Breathing Science is Life**.

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### **Executive Summary**

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#### **Program Overview**

This pilot NTM Journal Club program was delivered via Twitter, live webinar, and endured online. The multimedia NTM Journal Club sessions were developed and moderated by NJH faculty on a monthly basis, with downloadable article summaries that provided the key points of recently published articles in NTM and a group opinion developed by the National Jewish Health Infectious Disease Physicians Group. Each article summary issue is archived on a dedicated webpage. Every month, a thirtyminute live webinar led by expert NJH faculty provided a succinct article summary and engaged participants in academic and peer discussion. The recording of each live webinar is endured on VuMedi and made available for a year. A 30-minute structured Twitter chat based on the same article was also offered each month, providing another forum for live interaction with peers and expert faculty, as well as ongoing Tweet exchange for those that could not attend the live portion.

#### **Program Chairs**



Charles Daley, MD
Chief, Division of
Mycobacterial and
Respiratory Infections
National Jewish Health



### Shannon H. Kasperbauer, MD

Associate Professor,
Division of Mycobacterial
and Respiratory Infections
National Jewish Health

#### **Learning Objectives**

- Apply critical thinking for research analysis in the review of new data and guidelines in NTM.
- Utilize increased awareness and understanding of research, evidence and best practices to inform clinical practice in NTM.
- Support an online community of practitioners to share key insights, latest research, and treatment strategies for patients with NTM.

#### **Program Webpage**

Launch Date: August 24, 2021 End Date: August 27, 2022

**Activity Link:** 

https://www.nationaljewish.org/ntmjournalclub

#### **Target Audience & Accreditation**

Target Audience: Pulmonologists, Infectious Disease Physicians, Primary Care/Family Medicine Physicians, Physician Assistants, and Nurse Practitioners.

Accreditation: National Jewish Health is accredited by the Accreditation Council for Continuing Medical Education (ACCME) to provide continuing medical education for physicians. NJH designates each of the 24 live activities (12 live webinars, 12 live Twitter chats) for a maximum of 0.5 AMA PRA Category 1 Credit™.

### **Audience Generation**

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**JOURNAL CLUB** 

NJHealth.org/NTMJournalClub

2→Zoom Discussion: Oct. 26, 2:30 p.m. EST

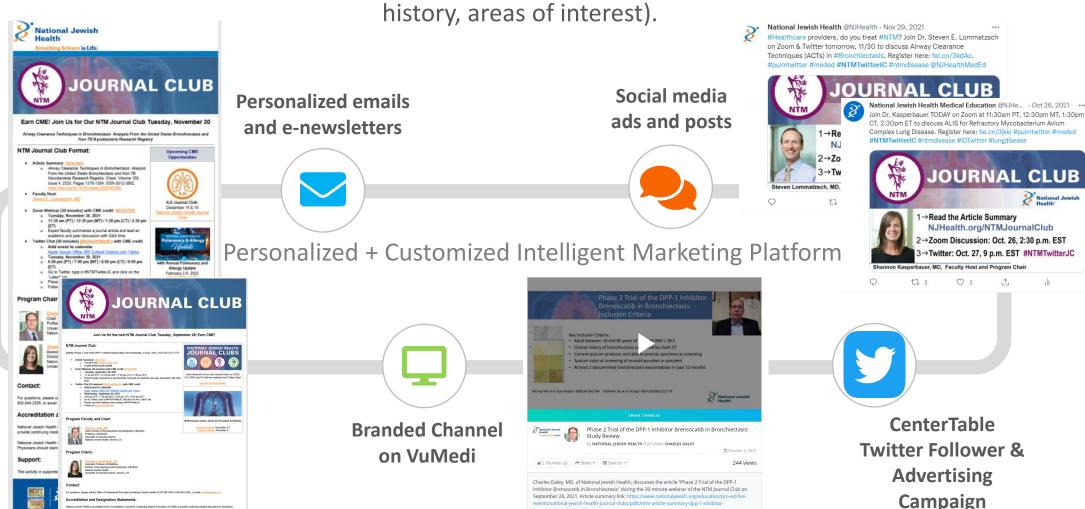
3→Twitter: Oct. 27, 9 p.m. EST #NTMTwitterJC

→ Read the Article Summary

0 2

Personalized targeting tools across numerous tactics reach HCPs by leveraging demographic data (such as location, profession, specialty) and behavioral data (such as learner participation

> brensocatib vol1 issue2.aspx 5 read less



### **Activity Format**

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## National Jewish Health Monthly NTM Journal Club Activities



### Article Summary

developed by NJH experts in NTM

endured on NJH Website



### Article Summary

sent to target
audience with
registration links to
Zoom and Twitter
Journal Club



### Live Webinar Journal Club via Zoom

30-minute discussion led by NJH NTM expert

(CME Credit)



Twitter Journal Club via Tweet Chat

(CME Credit)



Webinars Endured on Vu-Medi

12

article summaries developed

1,515

article summary downloads

276

webinar completers (548 registrants)

20,097 impressions

1,903 views on VuMedi

### Quantitative Educational Impact Summary



4%

N=55

22%

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Registrations

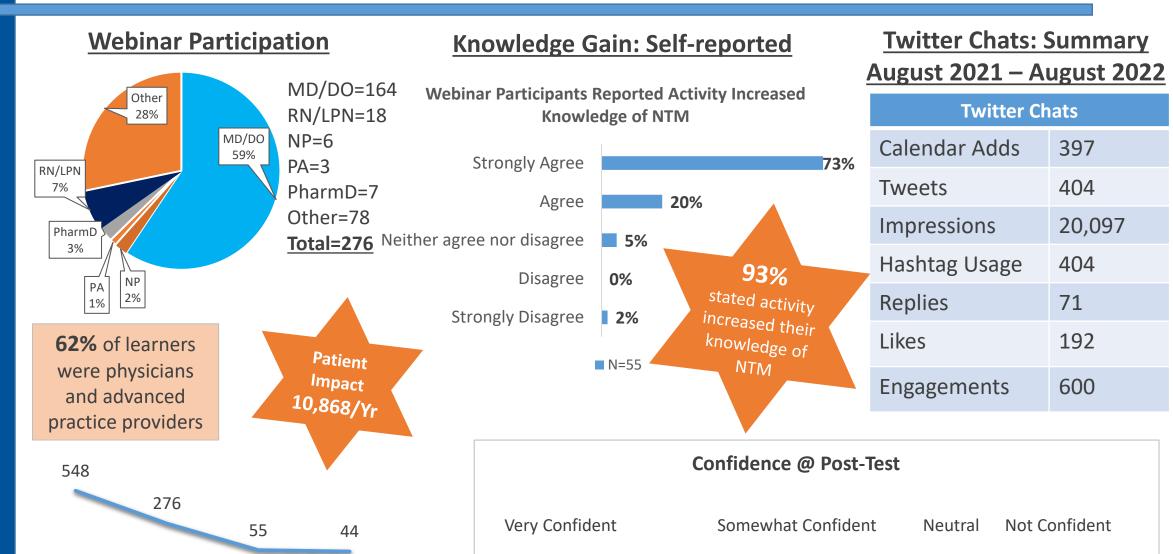
Webinar

Attendees

Evaluation

Respondents

Certs



38%

36%

### Qualitative Educational Impact Summary

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### **Patient Impact**

55

Evaluation respondents

Who see

209

**NTM Patients Weekly** 

Which translates to

10,868

**Patient Visits Annually** 

### **Educational Impact**

**Knowledge and Satisfaction: Self-reported** 

93% stated activity increased their knowledge of NTM [N=55]

**96%** indicated the learning objectives of the activity were met [N=55]

**82%** said the activity improved their ability to treat or manage patients [N=55]

#### **Practice Change**

91%

Reported intent to change their practice as a result of what they learned in the activity [N=45]

## Top Intended Practice Changes

- Incorporate omadacycline into the management of M. abscessus patients
- Emphasize the importance of airway clearance
- Implement the use of ALIS
- Implement screening tools to identify this disease

"The addition of the Twitter discussions is a great option!"
-NTM Journal Club Attendee

### Program Insights

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- In this pilot program, we found that most participants are not seeking credit for webinars or Twitter chats, though they are engaging with the content.
- Article downloads and endured video views are high, indicating high levels of participation in the educational content beyond the live activities.
- It appears not all Twitter chat participants are "active" in the live activity.
   However, based on data for engagements and likes, there are many viewers consuming the education presented without posting comments and actively contributing to the discussion.

"Our goal was to bring quality CME activities to colleagues and foster discussion over the latest studies and trials in nontuberculous mycobacteria. The Twitter Journal Club was a fun way to push the envelope and offer free open access medical education to an even broader audience. We were grateful for the opportunity to share in learning with colleagues from around the country and world."

Charles Daley, MD (NTM Journal Club Program Co-Chair)

### **Article Summary**

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August 2021 Article Summary: Amikacin liposome inhalation suspension for Treatment-Refractory lung disease caused by Mycobacterium avium complex (CONVERT). A prospective, open-label, randomized study. *Am J Repir Crit Care Med* 2018; 198: 1559-1569.

**View Here** 

**September 2021 Article Summary:** Phase 2 Trial of the DPP-1 Inhibitor Brensocatib in Bronchiectasis, *N Engl J Med* 2020; 383:2127-2137, DOI: 10.1056/NEJMoa2021713. View Here

October 2021 Article Summary: Amikacin Liposome Inhalation Suspension for Refractory Mycobacterium avium Complex Lung Disease: Sustainability and Durability of Culture Conversion and Safety of Long-term Exposure. Griffith DE, et al; CONVERT Study Group. Chest. 2021 Sep;160(3):831-842. DOI: 10.1016/j.chest.2021.03.070. View Here

November 2021 Article Summary: Airway Clearance Techniques in Bronchiectasis: Analysis From the United States Bronchiectasis and Non-TB Mycobacteria Research Registry. *Chest*, Volume 158, Issue 4, 2020, Pages 1376-1384, ISSN 0012-3692, https://doi.org/10.1016/j.chest.2020.06.050. View Here

January 2022 Article Summary: Preliminary, Real-world,
Multicenter Experience With Omadacycline for Mycobacterium
abscessus Infections. *Open Forum Infectious Diseases*, Volume 8,
Issue 2, February 2021, ofab002. View Here

### Article Summary

Final Report: August 2021 — August 2022



February 2022 Article Summary: Mycobacteriophageantibiotic therapy promotes enhanced clearance of drugresistant Mycobacterium abscessus. Matt D Johansen, Matthéo Alcaraz, Rebekah M Dedrick, Françoise Roquet-Banères, Claire Hamela, Graham F Hatfull, Laurent Kremer. s Model Mech. 2021 Sep 1;14(9):dmm049159. View Here

March 2022 Article Summary: Pilot study to test inhaled nitric oxide in cystic fibrosis patients with refractory Mycobacterium abscessus lung infection. *J Cyst Fibros*. 2020 Mar;19(2):225-231. View Here

April 2022 Article Summary: Mycobacterium abscessus biofilms have viscoelastic properties which may contribute to their recalcitrance in chronic pulmonary infections. *Sci Rep.* 2021 Mar 3;11(1):5020. doi: 10.1038/s41598-021-84525-x. PMID: 33658597. View Here

1515

total

article

downloads

May 2022 Article Summary: Genomic Analysis of a Hospital-Associated Outbreak of Mycobacterium abscessus: Implications on Transmission. *J Clin Microbiol*. 2022 Jan 19;60(1):e0154721. View Here

June 2022 Article Summary: Host and pathogen response to bacteriophage engineered against Mycobacterium abscessus lung infection. *Cell.* 2022 May 26;185(11):1860-1874.e12. doi: 10.1016/j.cell.2022.04.024. Epub 2022 May 13. PMID: 35568033. View Here

July 2022 Article Summary: Consensus management recommendations for less common non-tuberculous mycobacterial pulmonary diseases. Lancet Infect Dis. 2022 Jul;22(7):e178-e190. doi: 10.1016/S1473-3099(21)00586-7. Epub 2022 Jan 25. Erratum in: Lancet Infect Dis. 2022 Mar;22(3):e73. PMID: 35090639. View Here

August 2022 Article Summary: Time to Positive Culture Detection Predicts Mycobacterium avium Pulmonary Disease Severity and Treatment Initiation. *Ann Am Thorac Soc.* 2022 Jun;19(6):925-932. doi: 10.1513/AnnalsATS.202107-765OC. PMID: 34851813.

View Here

National Jewish Health

Webinar Date	Article Title	Faculty	Registrations	Attendance
August 24, 2021	Amikacin liposome inhalation suspension for Treatment-Refractory lung disease caused by Mycobacterium avium complex (CONVERT). A prospective, open-label, randomized study.	David Griffith, MD	43	24
September 28, 2021	Phase 2 Trial of the DPP-1 Inhibitor Brensocatib in Bronchiectasis	Charles Daley, MD	26	13
October 26, 2021	Amikacin Liposome Inhalation Suspension for Refractory Mycobacterium avium Complex Lung Disease: Sustainability and Durability of Culture Conversion and Safety of Long-term Exposure	Shannon Kasperbauer, MD	28	18
November 30, 2021	Airway Clearance Techniques in Bronchiectasis: Analysis From the United States Bronchiectasis and Non-TB Mycobacteria Research Registry	Steve Lommatzsch, MD	50	33
January 25, 2022	Preliminary, Real-world, Multicenter Experience With Omadacycline for Mycobacterium abscessus Infections	Jared Eddy, MD	61	34

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Webinar Date	Article Title	Faculty	Registrations	Attendance
February 22, 2022	Mycobacteriophage-antibiotic therapy promotes enhanced clearance of drug-resistant Mycobacterium abscessus	Michael Strong, MD	44	23
March 29, 2022	Pilot study to test inhaled nitric oxide in cystic fibrosis patients with refractory Mycobacterium abscessus lung infection	Jane E. Gross, MD, PhD	36	14
April 26, 2022	Mycobacterium abscessus biofilms have viscoelastic properties which may contribute to their recalcitrance in chronic pulmonary infections	Jennifer R. Honda, PhD	38	16
May 24, 2022	Genomic Analysis of a Hospital- Associated Outbreak of Mycobacterium abscessus: Implications on Transmission	Rebecca Davidson, PhD	38	14

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Webinar Date	Article Title	Faculty	Registrations	Attendance
June 28, 2022	Host and pathogen response to bacteriophage engineered against Mycobacterium abscessus lung infection	Jerry Nick, MD	52	23
July 26, 2022	Consensus management recommendations for less common non-tuberculous mycobacterial pulmonary diseases	Charles Daley, Md	73	36
August 23, 2022	Time to Positive Culture Detection Predicts Mycobacterium avium Pulmonary Disease Severity and Treatment Initiation	Shannon Kasperbauer, MD	59	28
	Total August 2021 – August 2022			276



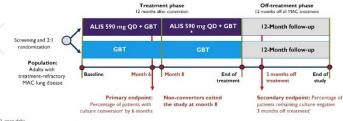
Final Report: August 2021 – August 2022



August 2021

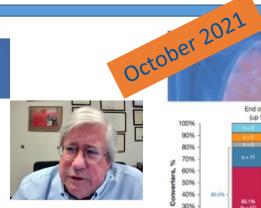
#### **CONVERT Study Design**

Converters remained in the study after month 8

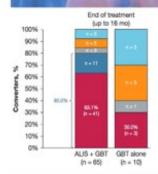


\*Culture conversion was defined as 3 consecutive monthly negative sputum cultures by month 5 Prespecified durability analysis: durability of culture conversion 3 months off all MAC treatment for patients who completed 12 months of treatment from the first negative culture that defined conversion.

Griffith DE, et al. Am J Respir Crit Care Med. 2018;198:1559-1569



#### Results

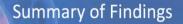












- · Brensocatib met both primary and secondary outcomes
- · Primary: prolonged time to first exacerbation c/w placebo
- Secondary: lower frequency of exacerbations, risk of exacerbations was approximately 40% lower than with placebo
- There was no significant change in ppFEV1 or Respiratory Symptom domain of the QOL-B bronchiectasis questionnaire
- Overall, brensocatib was well tolerated
  - · cough and dyspnea were more common in those who received brensocatib
  - · skin and dental events, both AEs of special interest, were more common with brensocatib
- Brensocatib is currently being evaluated in a Phase 3 randomized, placebocontrolled study (ASPEN) National Jewish

Chalmers JD, et al. N Engl J Med 220;383:2127-37



### Demographic Data

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Variable	Data Available, No.	Overall Sample (N = 905)	Continuous Use of Airway Clearance at Baseline and Follow Up (n = 226; 25%)	Intermittent Use of Airway Clearance at Baseline and Follow Up (n = 351; 39%)	No Use of Airway Clearance at Baseline and Follow Up (n = 328; 36%)	P Value
Exacerbations in the past 2 y, No. (%):	772	-	-	-		< .0001
0		254 (33)	38 (20)	89 (31)	127 (43)	
1		151 (20)	44 (23)	58 (20)	49 (17)	
2		135 (18)	31 (17)	57 (20)	47 (16)	
23		232 (30)	75 (40)	83 (29)	74 (25)	
Hospitalized for pulmonary illness or exacerbation in the past 2 y, No. (%)	864	228 (26)	70 (32)	90 (27)	68 (22)	.001
Non-tuberculous mycobacteria at baseline, No. (%)	805	255 (32)	64 (30)	104 (33)	87 (32)	.674
Pseudomonas aeruginosa, No. (%)	717	293 (41)	95 (47)	117 (40)	81 (36)	.021





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January 2022

#### Results (2)



7/12 isolates underwent subspeciation

Abscessus (6/7)

February 2024

- Massiliense (1/7)
- · Functional erm gene in 6/9
- 10/12 had antibiotics prior to omadacycline: median 4.7 months (IQR 3.4-12.7) [no data 1 patient]
- · 6/9 had positive cultures when omadacycline was initiated [missing data]
- Only 1 patient omadacycline MICs but 11/12 reported tigecyclin e) 4.3
- All patients with 2 or more companion drugs: 8/12 amikacin, 5/12 imipenem, 5/12 linezolid/tedizolid, 12 clofazimine, 2/12 tigecycline

- Median duration of omadacycline 6.2 months (IQR 4.2-11.0) [all oral therapy]
- Median duration of follow-up 5.1 months (IQR 3.4-7.2)
- Clinical success in 9/12 (75%)
- · Why omadacycline was used:
  - · Resistance to previous antibiotics (8/12)
  - Previous antibiotic failure (6/12)
  - Ease of administration (6/12)
  - · Oral bioavailability (6/12)
- · 3 adverse effects (all managed with drug retained)
  - · Nausea/vomiting/diarrhea
  - · Creatinine increase >0.5 mg/dL
  - · AST/ALT >3x upper limit normal



### Results (2)

#### iNO Safety:

March 2022

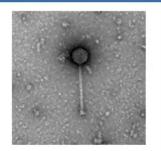
- 0 iNO-related serious adverse events (SAEs).
- 25 adverse events (AEs).
- All iNO-related AEs were minor, transient, and self resolved.

Event	n (pts) / n (event)	Comments
Common cold	1/1	
Pulmonary exacerbation	5/6	
Dizziness	1/2	Possibly NO-related, resolved
Dry mouth	1/1	Probably NO-related <sup>a</sup> , resolved
Hemoptysis	3/4	Possibly NO-related ( $n = 1$ ), minor, resolved
Musculoskeletal pain	1/1	
Pain in extremity	1/2	
Headache	1/1	
Vomiting	2/2	
Fever	2/3	
MetHb elevation	1/1	Probably NO-related (7.1%)
Papilledema (blurred vision)	1/1	transient, resolved, SAE
Patients with probable/possible	3	minor, resolved



#### A Phage Named Muddy

Det	ailed Information for Phage Muddy	
Discovery Information		
Isolation Host	Mycobacterium smegmatis mc²155	
Found By	Lilli Holst	
Year Found	2010	
Location Found	Durban, South Africa	
Finding Institution	University of Kwazulu-Natal	
Program	Mycobacterial Genetics Course, Durban, South Africa	
From enriched soil sample?	Yes	
Isolation Temperature	Not entered	
GPS Coordinates	29.8587 S, 31.0218 E Map	
	Muddy was the 1400th phage entered into the Mycobacteriophage DataBase!	
Discovery Notes	This sample was scraped from the underside of a partially decomposed aubergine and was therefore largely decomposed vegetable matter. The sample was dark, moist, and worms and other insects were present.	

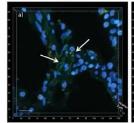




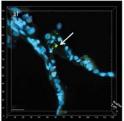
#### M. abscessus lung biofilms

1st report of M. abscessus biofilms in a human lung

M. abscessus biofilms in the thickened alveolar walls of explanted lung tissue from a pwCF.



Qvist, et al., ERS J, 2015





Fennely, et al., AJRCCM, 2016



phagesdb org

**National Jewish** 



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#### Whole Genome Sequencing (WGS) of NTM isolates

Bryant et al 2016; Science. 354(6313)

Population genomics studies of M. abscessus from persons with CF (pwCF) show highly similar genetic clones

Outbreak clusters in a single CF center (UK) \*Bryant et al 2013; Lancet. 381(9877)

Genetic similarity defined in terms of single nucleotide polymorphisms (SNPs)

- <20 SNPs "probable" transmission
- 20-38 SNPs "possible" transmission
- Person-to-person transmission\*
- Direct or indirect?
- Clones originated from the same country and different countries



Consensus management recommendations for less common non-tuberculous mycobacterial pulmonary diseases

Christoph Lange, Erik C Bottger, Emmanuelle Cambau, David E Griffith, Lorenzo Guglielmetti, Jakko van Ingen, Shandra L Knight, heodore K.Marras, Kenneth N. Olivier, Miguel Santin, Jason E. Stout, Enrico Tortoli, Dirk Wagner, Kevin Winthrop, Charles L. Daley, on behalf of the opert panel group for management recommendations in non-tuberculous mycobacterial palmonary diseases

#### **Today's Article** Presented by: Charles L. Daley, MD

Consensus Management Recommendations for Less Common Non-tuberculous Mycobacterial Pulmonary Diseases Lange C, et al. Lancet Infect Dis 2022;22.e178-90



Dreambing Science is \$150.

June 2022

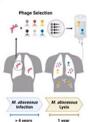
- 26 y/o male with Cystic Fibrosis
- · M. abscessus in 2016 · Enrolled in the PREDICT Trial
- · Met diagnostic criteria in early 2017
- · Enrolled in the PATIENCE Trial Refractory to antibiotic treatment
- · Two effective phages identified
- Initiated on phage therapy Sept 2020
- Culture conversion January 2021
- · Lung Transplant October 2021
- · Completed phage (and antibiotics)







Host and pathogen response to bacteriophage engineered against Mycobacterium abscessus lunc







#### Methods

- Retrospective cohort study
  - Patients seen at the Toronto Western Hospital NTM clinic between January 1, 2015 through December 21, 2019.
- Two positive sputa for M. avium
  - · Either expectorated or induced
  - "index" sputum (culture + for M. avium without treatment and recent CT)
  - . There was no time limit between the two sputa and only the index isolate was required to meet criteria for study inclusion.
  - If multiple sputa met criteria for the index, only the earliest sample was chosen.
- No active or prior treatment within the last six months.
- Patients had computed tomographic (CT) imaging of the chest within six months of their index sputum result. **National Jewish** Health'

### Level (1) Outcomes: NTM Journal Club Webpage



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### **NTM Journal Club Webpage**

https://www.nationaljewish.org/ntmjournalclub

Page Views 7,288

Unique Page Views 3,067



Article
Downloads

1,515

\*Data from 7/29/21 - 9/7/2022

National Jewish
Health Journal Clubs

**COPD Journal Club** 

**ILD Journal Club** 

**NTM Journal Club** 

PH Journal Club

How to Participate in a Journal Club Twitter Chat



#### Respiratory Medicine Updates: A Virtual Clinical Community

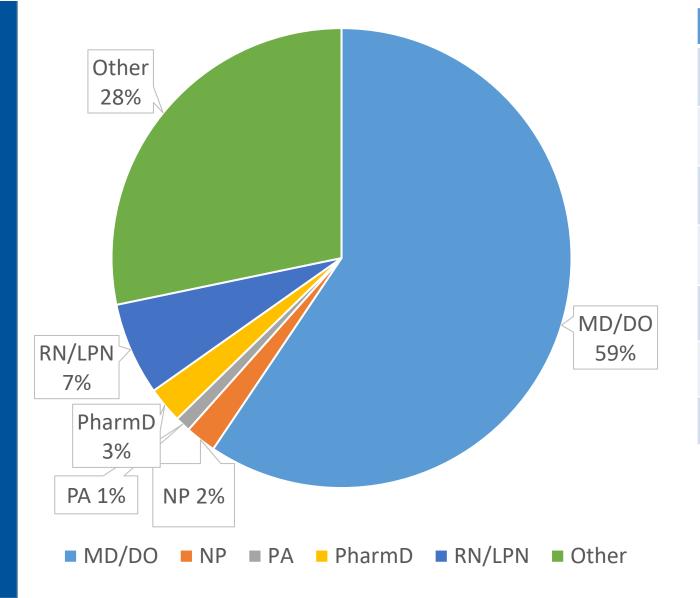
Our Next Session is August 23–24, 2022 • CME Credit for Zoom Webinar and Twitter Chat Participants

**Article:** Time to Positive Culture Detection Predicts *Mycobacterium avium* Pulmonary Disease Severity and Treatment Initiation. Ann Am Thorac Soc. 2022 Jun;19(6):925-932. doi: 10.1513/AnnalsATS.202107-765OC. PMID: 34851813.



### Level (1) Outcomes: Live Webinars Participation: By Degree



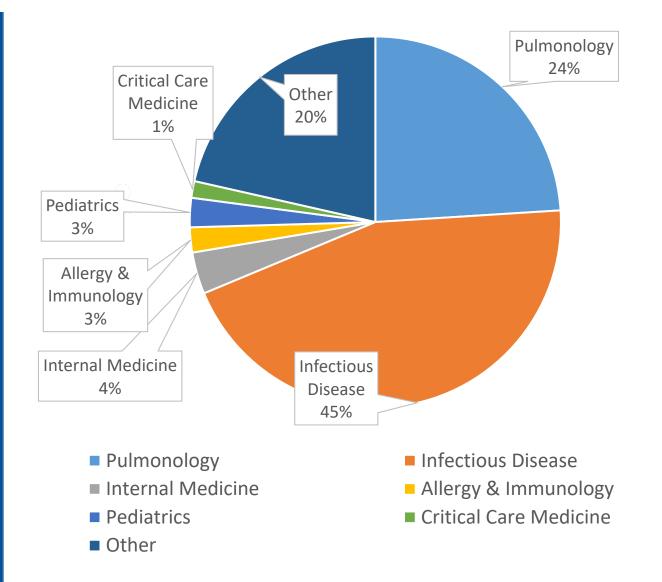


Degree	Completer #s
MD/DO	164
RN/LPN	18
NP	6
PA	3
PharmD	7
Other	78
<b>Total Completers</b>	276



### Level (1) Outcomes: Live Webinars Participation: By Specialty



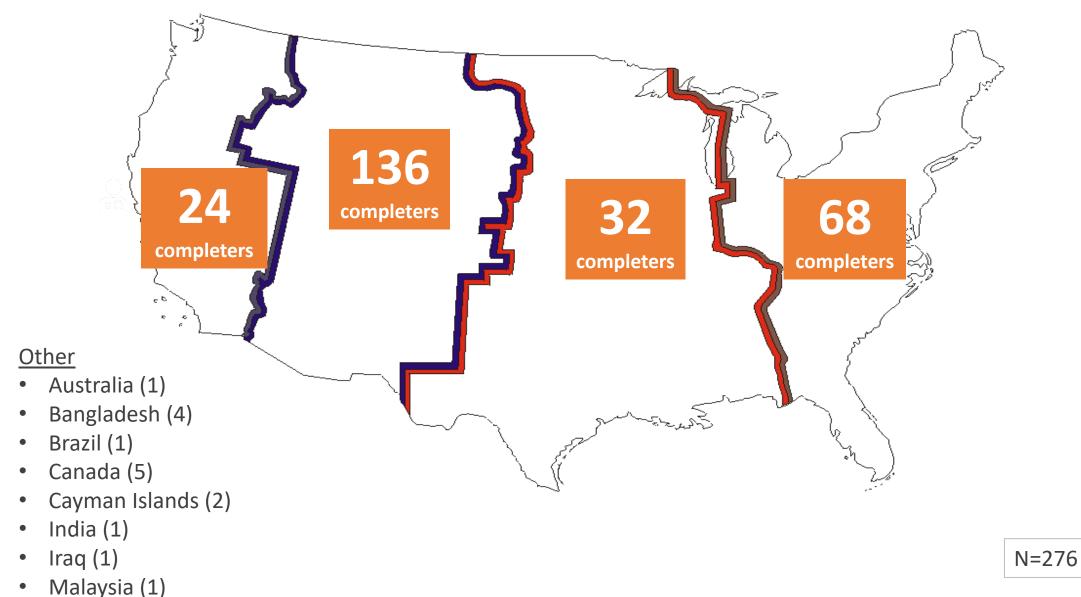


Specialty	Completer #s
Infectious Disease	123
Pulmonology	66
Internal Medicine	10
Allergy & Immunology	6
Pediatrics	7
Critical Care Medicine	4
Other	60
<b>Total Completers</b>	276



### Level (1) Outcomes: Live Webinars Participation



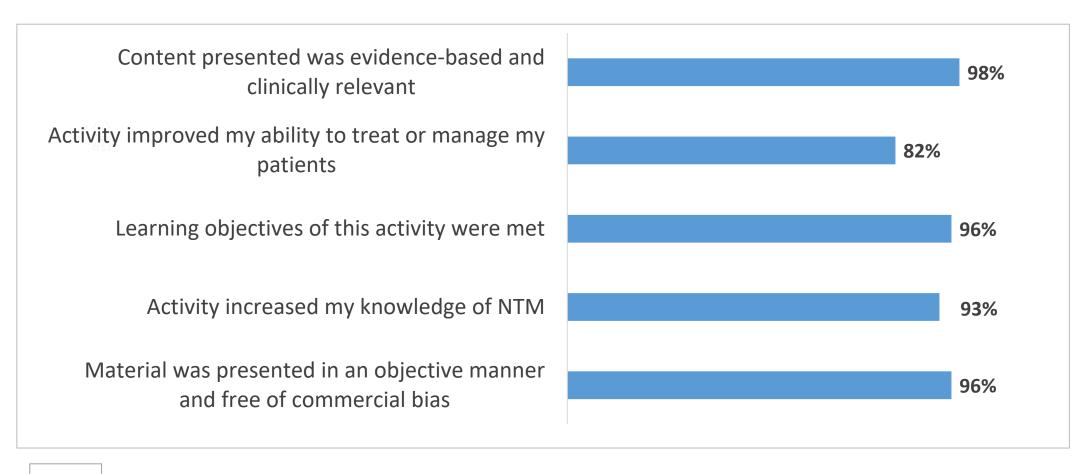




### Level (2&3) Outcomes: Live Webinars Satisfaction & Knowledge S National Jewish

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### **Evaluation respondents "Strongly Agree" or "Agree" that:**

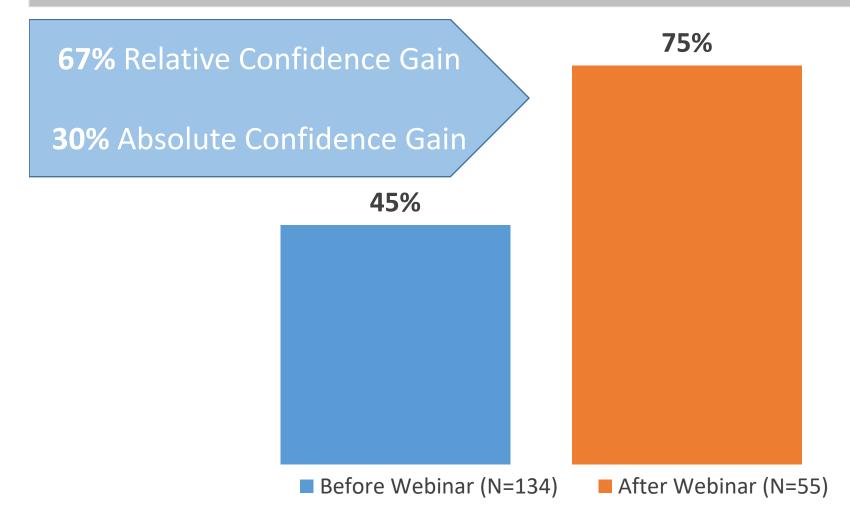






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Evaluation respondents report they are "very confident" to "somewhat confident" in their ability to integrate the findings of the research article into clinical practice:







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### An analysis of open-ended comments demonstrates the following changes completers intend to make:

Month	Article Title	Intended Changes
August 2021	ALIS for Treatment-Refractory Lung Disease Caused by MAC	<ul> <li>Implement the use of ALIS (3 responses)</li> <li>Utilize strategies to manage side effects</li> <li>Provide motivation to patients</li> </ul>
September 2021	Phase 2 Trial of the DPP-1 Inhibitor Brensocatib in Bronchiectasis	<ul> <li>Use strategies to identify these patients in practice</li> <li>Implement screening tools to identify this disease</li> </ul>
October 2021	ALIS for Refractory MAC Lung Disease: Sustainability and Durability of Culture Conversion & Safety of Long-term Exposure	<ul> <li>Use ALIS as a treatment for longer (2 responses)</li> <li>Utilize testing/screening to select patients who would benefit from this treatment</li> </ul>
November 2021	Airway Clearance Techniques in Bronchiectasis: Analysis from the US Bronchiectasis and Non-TB Mycobacteria Research Registry	<ul> <li>Emphasize and educate patients on the importance of airway clearance (5 responses)</li> <li>Use strategies to evaluate patients for symptoms of bronchiectasis</li> </ul>
January 2022	Multicenter Experience with Omadacycline for Mycobaterium abscessus Infections	<ul> <li>Utilize omadacycline as part of treatment strategy (5 responses)</li> </ul>
February 2022	Mycobacteriophage-antibiotic therapy promotes enhanced clearance of drugresistant Mycobacterium abscessus	<ul> <li>Evaluate CF patients for NTM</li> <li>Include phage therapy as first option when possible</li> <li>Apply animal model evidence in antibiotic selection</li> </ul>





Final Report: August 2021 – August 2022

### An analysis of open-ended comments demonstrates the following changes completers intend to make:

Month	Article Title	Intended Changes
March 2022	Pilot study to test inhaled nitric oxide in cystic fibrosis patients with refractory Mycobacterium abscessus lung infection	<ul> <li>Consider NO as a therapeutic modality for patients with CF or CFTR-related disorder</li> <li>Study this subject further</li> </ul>
April 2022	Mycobacterium abscessus biofilms have viscoelastic properties which may contribute to their recalcitrance in chronic pulmonary infections	Use a lower treatment threshold for Mabs patients
May 2022	Genomic Analysis of a Hospital-Associated Outbreak of Mycobacterium abscessus: Implications on Transmission	Be aware of the limitations of WGS in evaluating M. abscessus outbreaks
June 2022	Host and pathogen response to bacteriophage engineered against Mycobacterium abscessus lung infection	<ul> <li>Discuss article with our contracted ID specialist</li> <li>Consider submitting sputum to my refractory cases for phage consideration</li> </ul>





Final Report: August 2021 – August 2022

### An analysis of open-ended comments demonstrates the following changes completers intend to make:

An analysis of open-ended comments demonstrates the following changes completers intend to make:			
Month	Article Title		Intended Changes
July 2022		agement recommendations for on-tuberculous mycobacterial asses	<ul> <li>Empiric therapy based on species</li> <li>Reinforced current management. Also will consider use of inhaled tobramycin for M. chelonae pulmonary</li> <li>Increase knowledge of NTM to help with TB differential</li> </ul>
August 2022	Time to Positive Culture Detection Predicts Mycobacterium avium Pulmonary Disease Severity and Treatment Initiation		<ul> <li>Consider submitting sputum to my refractory cases for phage consideration.</li> <li>Discuss article with our contracted ID specialist</li> <li>Pay closer attention to the time to sputum culture positivity</li> <li>Ensure I evaluate the amount of airway clearance my</li> </ul>
9:	<b>1%</b> N=45	Evaluation respondents intend to make changes in practice as a result of the activity	<ul> <li>Keeping time to positivity in mind when looking at sputum cx for MAC</li> </ul>



Final Report: August 2021 – August 2022



August: ALIS for Treatment-Refractory
Lung Disease Caused by MAC
(CONVERT Study)



#### **Key Takeaways**

- Strong evidence for the use of ALIS
- Use ALIS for treatment-refractory disease (3 responses)



#### Questions

- I've had a couple of patients with dysphonia that resolved after cessation of ALIS for a few days and curiously did not recur when they resumed the medication. I cannot explain this...
- May not be related to trial, but wondering whether there is any data/evidence in using ALIS as a third drug along with macrolide and rifampin in ethambutol intolerant patients with nodular MAC?
- IV amikacin is given thrice weekly...do you think this might work just as well since amikacin is a concentration dependent antibiotic?

**September:** Phase 2 Trial of the DPP-1 Inhibitor Brensocatib in Bronchiectasis



#### **Questions**

- What were the reasons for more females in the trial?
- If brensocatib is put on the market, what steps will patients need to take to prevent the dental events?
- How do you see placement of this drug in the management of bronchiectasis? Will it be primary therapy or as a complement to airway clearance and other standard approaches?
- Was there any difference in outcomes between smokers and nonsmokers?



#### **Future Topics**

- Emerging therapies for NTM
- Phage Therapy
- Use of macrolides



Final Report: August 2021 — August 2022



October: ALIS for Refractory MAC Lung Disease: Sustainability & Durability of Culture Conversion & Safety of Long-term Exposure



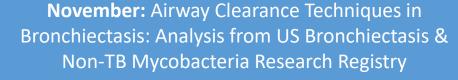
#### **Key Takeaways**

- ALIS has a durable and sustained response
- Safe and effective to give ALIS longer
- ALIS helps with culture conversion



#### Questions

- Why did the study randomize in a 2:1 ratio? Did the low number of non-ALIS patients possibly effect the outcomes?
- Does the fact that good outcomes were seen even for some patients who did not complete 12 months argue that we could shorten treatment duration?
- Was ototoxicity assessed objectively? If so, I assume there was no difference in this side effect between ALIS+GBT vs GBT?





#### **Key Takeaways**

- Emphasis on keeping up airway clearance over time
- Mucus clearance techniques
- ACT does not seem to have a global positive impact
- A patient's use of ACTs can tell you a lot about their disease course and symptoms



#### Questions

- Are you aware of any prospective studies evaluating the effect of ACT or pulmonary rehab on exacerbation rate / clinical decline in recently diagnosed NCFBE patients?
- Does ACT reduce pseudomonal airway colonization?
- What about prospective studies on mucolytic therapies and exacerbations in conjunction with ACT?



Final Report: August 2021 — August 2022



January: Preliminary, Real-world, Multicenter
Experience With Omadacycline for Mycobacterium
abscessus Infections



#### **Key Takeaways**

- Potential activity of omadacycline for M. abscessus
- Omadacycline is promising
- Small study, but effective and welltolerated in the follow-up period.
- Omadacycline has some data behind its use in M. abscessus and that PO omadacycline is an appropriate choice.



#### Questions

- Do you think it is necessary to test for omadacycline susceptibility or is tigecycline susceptibility adequate to presume susceptibility?
- Thoughts about dosing omadacycline at 15( mg bid vs 300 mg daily impact on PK/PD and tolerability?
- Any guidance on what range of tigecycline MIC would be acceptable to use omadacyline?



**February:** Mycobacteriophage-antibiotic therapy promotes enhanced clearance of drug-resistant Mycobacterium abscessus

#### **Key Takeaways**

- Phage therapy holds promise in the field of treating NTM infectious disease
- Tools for mycobacterial genetics
- Being informed how effective therapy is
- In the zebra fish model, phage therapy seemed to have some benefit though not enough to reverse mortality trends. There was an enhanced benefit in zebra fish with the CF mutation, though the mechanism of this benefit was unclear to me.



#### **Future Topics**

- More skin diseases
- Prevention for those infections
- A journal club about using beta lactam/beta lactamase inhibitors in patients with MAb would be helpful. Other articles on treatment resistant NTM and also surgical journal clubs.
- Role of PK monitoring



Final Report: August 2021 – August 2022



March: Pilot study to test inhaled nitric oxide in cystic fibrosis patients with refractory Mycobacterium abscessus lung infection



#### **Key Takeaways**

- Nitric oxide may be a potential future drug for NTM
- NO may be an additional therapy for my NTM patients
- NO as an NTM treatment is a topic that is new for me and I have to read more about it
- Importance of having a larger sample size during studies



#### **Questions/Comments**

- What about airway clearance as a confounding factor?
- Early phase study recruiting in Australia using the home NO system (LungFit) for treatment of NTM in CF and non CF patients
- NO was being given for its direct effect on the NTM

**April:** Mycobacterium abscessus biofilms have viscoelastic properties which may contribute to their recalcitrance in chronic pulmonary infections



#### **Key Takeaways**

 The reasons that airway clearance may be limited for Mabs patients

May: Genomic Analysis of a Hospital-Associated
Outbreak of Mycobacterium abscessus: Implications
on Transmission

No qualitative data available for May



Final Report: August 2021 — August 2022



June: Host and pathogen response to bacteriophage engineered against Mycobacterium abscessus lung infection

#### **Key Takeaways**

- Phage therapy can be effective to help treat M. abscessus in some patients.
- Increased knowledge of TB differentials/NTM
- Phage may have a significant use in the treatment of bacterial disease, but MUCH more data is needed
- With appropriate treatment we can help clear patients of NTM and allow for better quality of care.
- Usefulness of bacteriophage

#### **Questions/Comments**

 Did you find any mutations in the post phage treatment persisters that may be associated to this resistant phenotype?



July: Consensus management recommendations for less common non-tuberculous mycobacterial pulmonary diseases

#### **Key Takeaways**

- Education about this published guideline
- Not much published research on NTMs, rely on expert consensus.
- The identification for treatment pathways is not always necessary

#### **Questions/Comments**

- For rapid growers, do you suggest omadacycline at any point in treatment?
- Is the mechanism for synergy between clofazimine and macrolide known?
- I have several patients treated for pulmonary MAC who have M. lentiflavum on surveillance sputum cultures. Is this likely an environmental phenomena? They are asymptomatic.
- For dosing ranges, for clofazimine and doxycycline how do you determine the dose? Body weight? Or when to use clofazimine 100 vs 200?
- Any experience using inhaled tobramycin for m. cholonae pulmonary infection?





Final Report: August 2021 — August 2022



**August:** Time to Positive Culture Detection Predicts Mycobacterium avium Pulmonary Disease Severity and Treatment Initiation



#### **Key Takeaways**

- Time to sputum culture positivity is important
- May be a good tool for the future
- Time to positivity may be helpful to look at, however it is hard to base any decisions off of it



#### **Questions/Comments**

- What are the main environmental sources of MAC?
- Does that include hot tubs and swimming pools? Does chlorine kill MAC?
- How effective are surface disinfectants like peroxide?
- Any extra tools to help make decisions?
- What group (mild/moderate) would you pick to study?
- Did you do a comparison with Jacko's (Netherlands) article to this one?
- Comment on airway clearance on time to positivity?

### Outcomes: VuMedi Webinar Recordings



Final Report: August 2021 — August 2022

#### **National Jewish Health VuMedi Channel:**

https://www.vumedi.com/channel/national-jewish-health/tab/journal-club/

Month	Impressions	Video Page Views	Unique Page Viewers	Starts	Unique Starts
August	2917	284	203	118	101
September	2260	257	177	132	102
October	2556	131	99	59	52
November	2443	284	188	155	116
January	1333	139	102	61	53
February	4986	150	104	73	52
March	1966	105	90	28	23
April	April 1795		84	31	25
May	4746	113	92	36	32
June	June 1179 July 342		70	40	32
July			161	103	82
August	156	8	5	5	4
Total	26,679	1,903	1,375	841	674

Data from 8.21.21 to 9.12.22

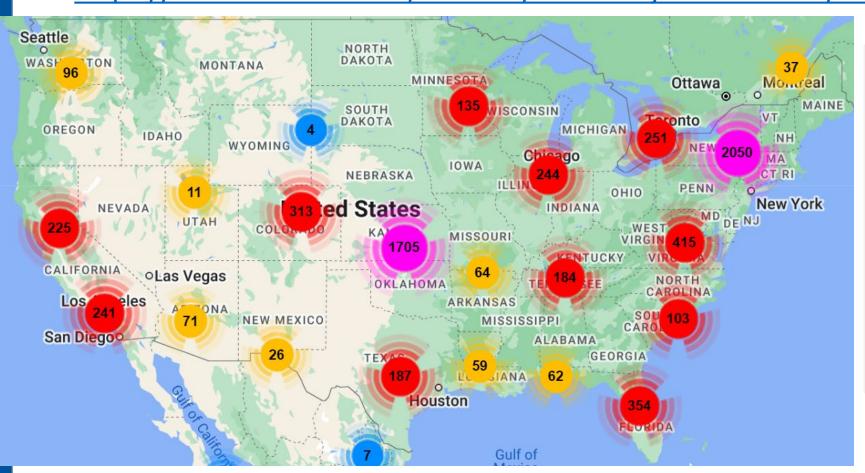
### Outcomes: VuMedi Webinar Recordings



Final Report: August 2021 — August 2022

#### **National Jewish Health VuMedi Channel:**

https://www.vumedi.com/channel/national-jewish-health/tab/journal-club/



#### **About VuMedi**

- Nearly 14,000 registered pulmonary specialists and over 96,000 primary care physicians (over 600,000 total)
- Distribution of video content to reach a large physician audience
- Analytics dashboard that shows video views, impressions, geolocation of viewers

Note: Heat map encompasses all 4 NJH Journal Clubs from 8/1/21-9/12/22



### National Jewish Health Medical Education Twitter

National Jewish Health

Final Report: August 2021 – August 2022

### Medical Education for Health Care Providers





Edit profile

#### **National Jewish Health Medical Education**

@NJHealthMedEd

The leading respiratory hospital provides continuing medical education, research insights, and best practices for patient care. | For patients, follow @NJHealth

O Denver, Colorado O njhealth.org/CME III Joined June 2021

296 Following 461 Followers



# Twitter Handle: @NJHealthMedEd

## Hashtag: #NTMTwitterJC



### Executive Summary – Twitter Chats

National Jewish Health

Twitter Date	Article Title	Faculty	Calendar Adds	Tweets	Retweets	Likes	Hashtag Usage	Impressions	Engagements
August 25, 2021	ALIS for Treatment-Refractory lung disease caused by MAC (CONVERT). A prospective, open-label, randomized study.	David Griffith, MD	45	33	1	25	33	3,302	81
September 29, 2021	Phase 2 Trial of the DPP-1 Inhibitor Brensocatib in Bronchiectasis.	Charles Daley, MD	24	40	6	38	40	1,846	57
October 27, 2021	ALIS for Refractory MAC Lung Disease: Sustainability and Durability of Culture Conversion and Safety of Long- term Exposure.	Shannon Kasperbauer, MD	56	36	1	0	36	1,430	33
November 30, 2021	Airway Clearance Techniques in Bronchiectasis: Analysis From the United States Bronchiectasis and Non-TB Mycobacteria Research Registry.	Steve Lommatzsch, MD	29	38	1	42	38	2,500	90
January 26, 2022	Preliminary, Real-world, Multicenter Experience With Omadacycline for Mycobacterium abscessus Infections.	Jared Eddy, MD	16	34	0	2	34	924	36



### Executive Summary – Twitter Chats



Twitter Date	Article Title	Faculty	Calendar Adds	Tweets	Retweets	Likes	Hashtag Usage	Impressions	Engagements
February 23, 2022	Mycobacteriophage- antibiotic therapy promotes enhanced clearance of drug- resistant Mycobacterium abscessus.	Michael Strong, MD	14	52	5	6	52	979	24
March 30, 2022	Pilot study to test inhaled nitric oxide in cystic fibrosis patients with refractory Mycobacterium abscessus lung infection.	Jane E. Gross, MD, PhD	46	21	6	8	21	1,544	52
April 27, 2022	Mycobacterium abscessus biofilms have viscoelastic properties which may contribute to their recalcitrance in chronic pulmonary infections.	Jennifer R. Honda, PhD	13	29	4	3	29	1,298	35
May 25, 2022	Genomic Analysis of a Hospital-Associated Outbreak of Mycobacterium abscessus: Implications on Transmission.	Rebecca Davidson, PhD	114	30	4	26	30	2,680	67
June 29, 2022	Host and pathogen response to bacteriophage engineered against Mycobacterium abscessus lung infection.	Jerry Nick, MD	8	31	5	2	31	1293	38



# Executive Summary – Twitter Chats



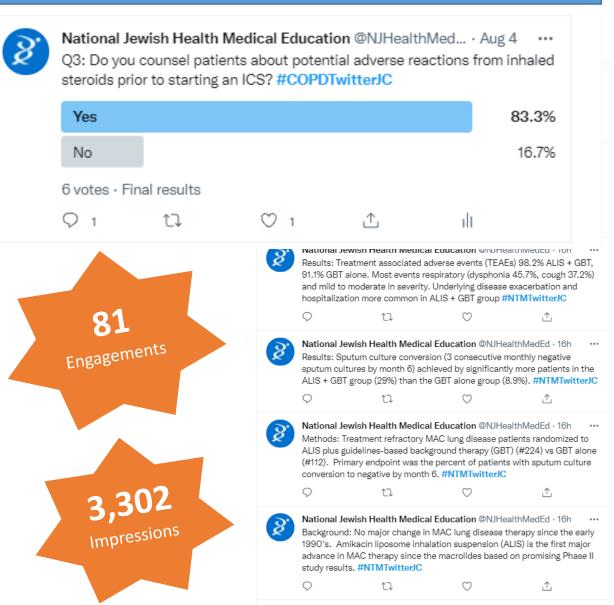
Twitter Date	Article Title	Faculty	Calendar Adds	Tweets	Retweets	Likes	Hashtag Usage	Impressions	Engagements
July 27, 2022	Consensus management recommendations for less common non-tuberculous mycobacterial pulmonary diseases.	Charles Daley, MD	20	29	2	33	29	1,489	67
August 24, 2022	Time to Positive Culture Detection Predicts Mycobacterium avium Pulmonary Disease Severity and Treatment Initiation.	Shannon Kasperbauer, MD	12	31	2	7	31	812	20
	Total August 2021 – A	August 2022	397	404	37	192	404	20,097	600



# NTM Journal Club: <u>August Twitter Highlights</u>









# NTM Journal Club: <u>September Twitter Highlights</u>



How	ional Jewish Health Medical Education © v important do you consider spirometry wh nosis? #COPDTwitterJC			spiron	you make a di netry improves nte? #COPDT\	s to norma					o ire confirmed in we may have o	our first drug t	
Ve	ery important	80%		Yes				50%		of bronchiects	ISIS! #INTIVITWI	ttenc	
So	omewhat important	20%		No				50%					
Ne	eutral	0%		4 votes ·	Final results						0	_	
No	ot important	0%		7:24 PM ·	Sep 8, 2021 · Twitte	r Web App			38				100
5 vo	otes · Final results			View 1	weet activity						1	ノム	
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S	Sensitivity		50%		Replying to @NJHealt For those who answe discussion? #COPDT	red, could part	icipants share why	or why not for the		-	3	W	
S	Specificity		50%		D 13	WILLEIDC	Ť	ılı				3	
4 v	votes · Final results			8	Add another Tweet					0:00 164 views	MY HARPY	DANCE	- 0
Q4:	tional Jewish Health Medical Education of How would you diagnose a patient with elemetry? Type your response starting with	mphysema and normal			ake Woodrow @Jake Replying to @NJHealt A3 This is a tough que airways disease is rev	t <mark>hMedEd</mark> estion but I thi	nk if spirometry be			7:21 PM · Sep 29, 20:		Ø	<u>ث</u>
A4 1 dem patie	e Woodrow @JakeWoodrow3 · Sep 8 There is controversy here but in my opinion nonstrable airflow limitation is not COPD. I lents are not enrolled in COPD trials so I do rapies or not. #COPDTwitterJC	Important because these			Patricia George, MD A3. Question about the	his: what if the	y had abnormal sp	rometry and DLCO, bnml, and they have		Replying to 6 Exciting! Wh #INTMTwitte	tì	ose to use in the cum	rent phase 3 stud
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Patr	ricia George, MD @PGeorgeMD · Sep 8		) <i>I</i>			tì	$\bigcirc$	$\triangle$		Q	ti	♡ 1	±.
	olying to @JakeWoodrow3 and @NJHealthN this makes sense. I appreciate your though	MedEd Enga	agement!	S	ake Woodrow @Jake have had patients lik	ke this and I do	n't think we should			Replying to 6 So exciting!	Debb63 - Oct 2 RCLDaleyMD This is a common dise	ase and I'm sure will	be more so in the
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Repl	ricia George, MD @PGeorgeMD · Sep 8 olying to @NJHealthMedEd unk you for an informative #COPDtwitterJu	C, @JakeWoodrow3 and			Patricia George, MD Thank you!	@PGeorgeMD	- Sep 8	•••		Replying to 6	th bronchiectasis @bo RCLDaleyMD sed Dr. Daley! I didn't o		
@NII	JHealthMedEd			(	⊋ t	i,	$\odot$	<b></b>		0	t3	O 1	1

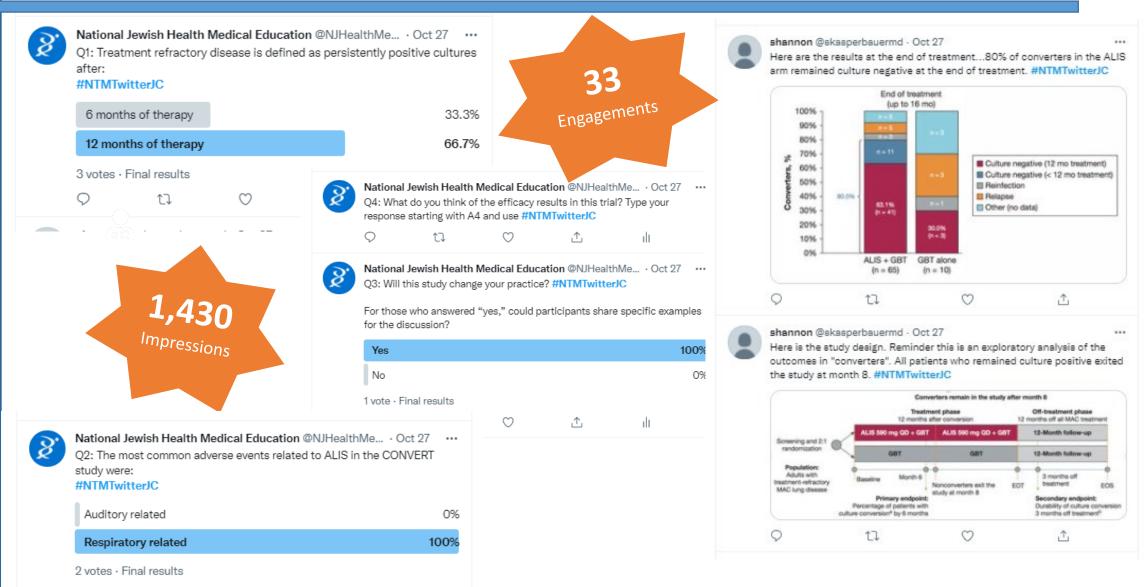


# NTM Journal Club: October Twitter Highlights



Final Report: August 2021 – August 2022

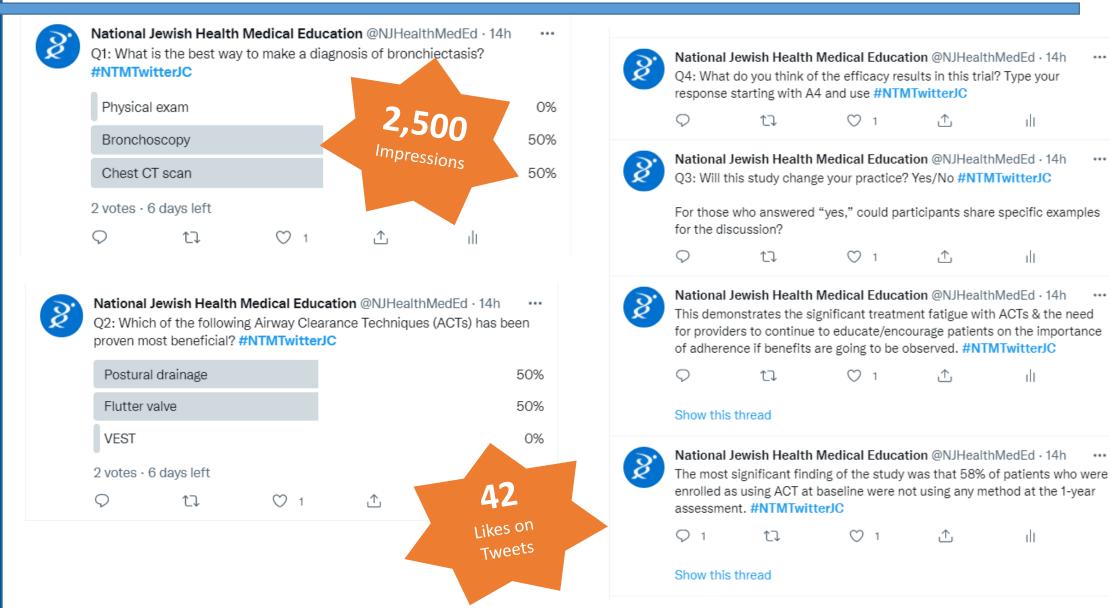
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# NTM Journal Club: November Twitter Highlights

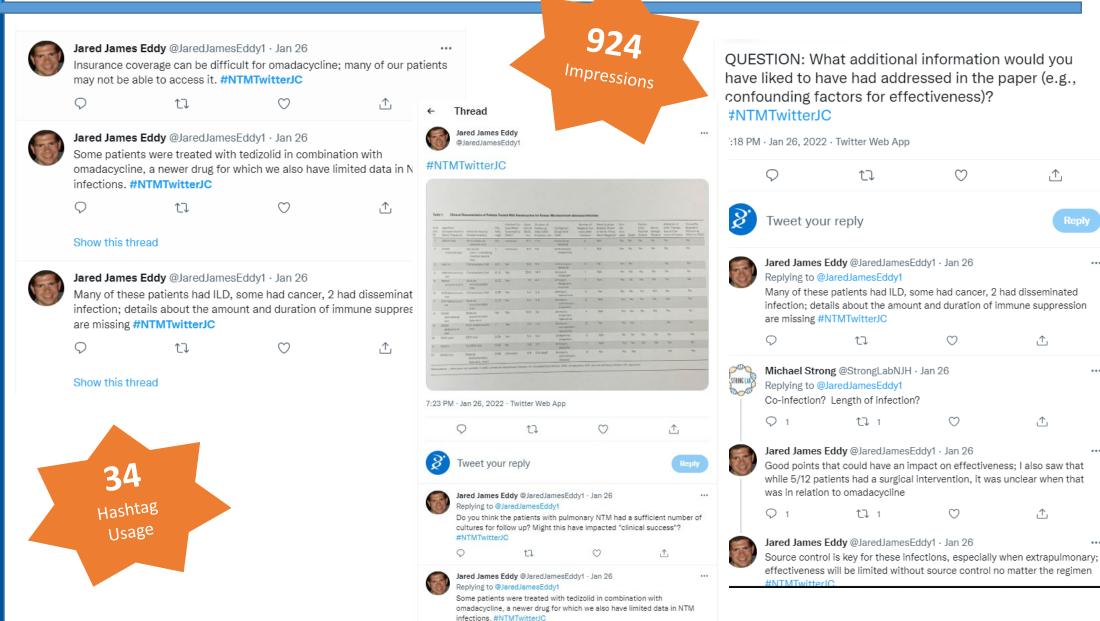






NTM Journal Club: <u>January Twitter Highlights</u>

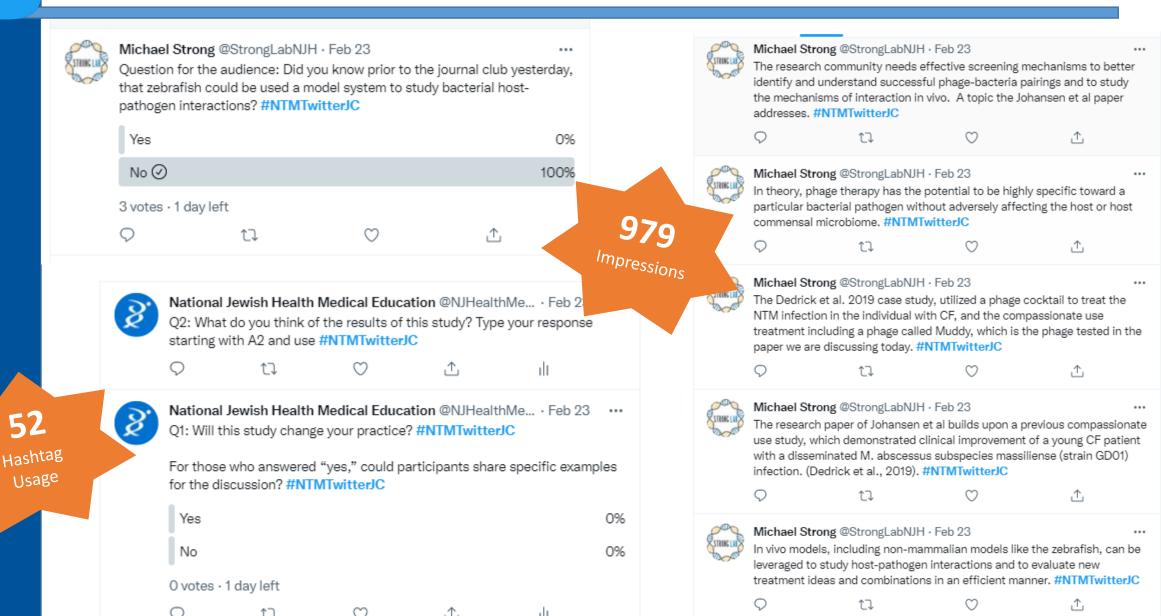






# NTM Journal Club: February Twitter Highlights



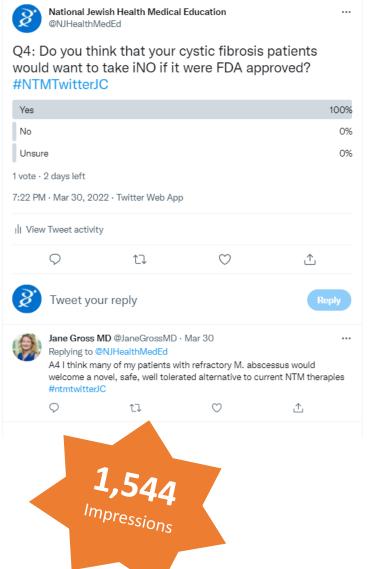




## NTM Journal Club: March Twitter Highlights









# NTM Journal Club: <u>April Twitter Highlights</u>

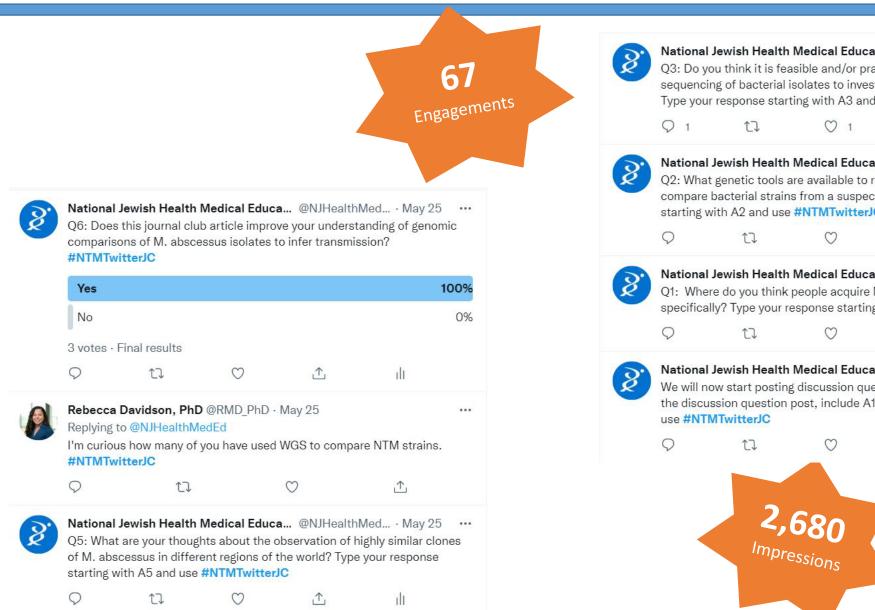
National Jewish Health





### NTM Journal Club: May Twitter Highlights

**National Jewish** 

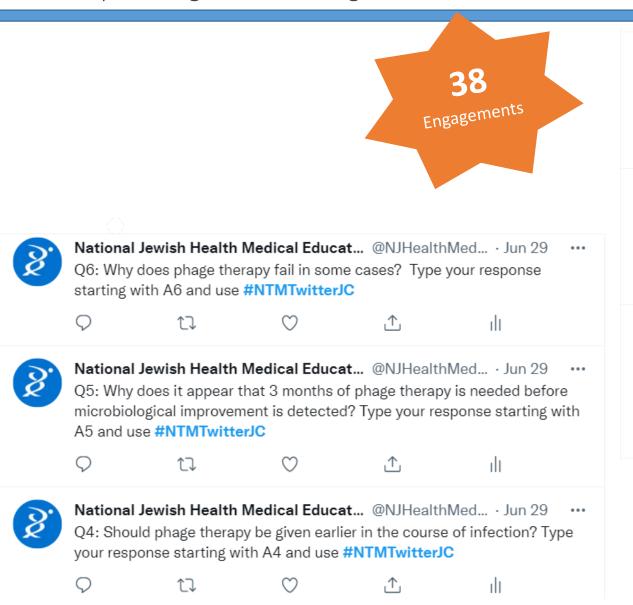


8.	Q3: Do you to sequencing o	hink it is feasibl of bacterial isola	e and/or practi ates to investiga	@NJHealthMed cal to use whole ate potential out e #NTMTwitter	genome breaks of NTM?
	Q 1	<b>↑</b>	♡ 1	$\triangle$	ili
8.	Q2: What ger compare bac	netic tools are a	available to rese om a suspected	@NJHealthMed. earchers or clinic outbreak? Type	cal labs to
	$\Diamond$	tì	$\bigcirc$	$\triangle$	ılt
8	Q1: Where d	o you think peo	ple acquire NTI	@NJHealthMed. M infections and ith A1 and use #	M. abscessus
	$\Diamond$		$\bigcirc$	$\triangle$	ilt
8	We will now s	start posting dis on question pos	cussion questi	@NJHealthMed. ons. To respond r Q1, A2 for Q2 a	, reply directly to
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# NTM Journal Club: June Twitter Highlights

National Jewish Health



8	Control of M. year of treatm there was no	abscessus allov ent. The explan evidence of pos	ved for a succe ted lungs did n t-transplant inf	@NJHealthMed. ssful lung transpot culture M. absection. Phage and the street and	olant after a scessus and nd antibiotic
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8.	Anti-phage ne failure. In this gradually agai	utralizing antibo subject, antipha	odies are a pote age-neutralizin age, indicating	@NJHealthMed. ential cause of p g antibodies tite different within- TwitterJC	hage therapy ers developed
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8.	Culture-indep with standard	endent markers airway cultures significant decr	were used to t These marker	@NJHealthMed. rack the infections s strongly suppo cessus burden w	on combined orted the
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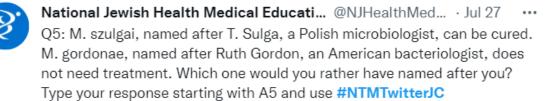


# NTM Journal Club: July Twitter Highlights

National Jewish Health

Final Report: August 2021 – August 2022





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National Jewish Health Medical Educati... @NJHealthMed... · Jul 27

Q4: When isolated, M. simiae is usually nonpathogenic. So why is it so difficult to treat? Type your response starting with A4 and use

#NTMTwitterJC

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National Jewish Health Medical Educati... @NJHealthMed... · Jul 27 ··· Q3: Has anyone seen a case of M. genavense pulmonary disease? Type your response starting with A3 and use #NTMTwitterJC

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National Jewish Health Medical Educati... @NJHealthMed... · Jul 27 ··· For M. chelonae, suggest ≥ 2-3 drugs with imipenem, tobramycin plus azithromycin, clofazimine, moxifloxacin, or linezolid. Same for M. fortuitum except amikacin instead of tobramycin, no azithromycin and additional drugs include cefoxitin, trim/sulfa and doxycycline#NTMTwitterJC

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National Jewish Health Medical Educati... @NJHealthMed... · Jul 27 ···

This year, members of the expert panel published a consensus statement on how to manage less common NTM. Systematic reviews were performed on seven species including 2 rapid growers and 5 slow growers and the results led to consensus treatment recommendations
#NTMTwitterJC

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National Jewish Health Medical Educati... @NJHealthMed... · Jul 27 ·· In 2020 the NTM tx guidelines were revised by a multi-society panel of experts that decided to focus on the most common NTM to cause pulmonary disease in adults without cystic fibrosis or immunosuppression. They included MAC, M. kansasii, M. xenopi, and M. abscessus #NTMTwitterJC

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# NTM Journal Club: August Twitter Highlights



Final Report: August 2021 - August 2022





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National Jewish Health Medical Educat... @NJHealthMed... · Aug 24 · · · · Q1: In this study did a shorter time to positivity predict M. avium pulmonary disease?

Type your response starting with A1 and use #NTMTwitterJC

National Jewish Health Medical Educat... @NJHealthMed... · Aug 24 · · · · We will now start posting discussion questions. To respond, reply directly to the discussion question post, include A1 for Q1, A2 for Q2 and so on, and use #NTMTwitterJC



	<10 Days (n = 36)	>10 Days (n = 89)	P Value
ase presence	29 (80.6)	52 (58.4)	0.02
smear positive	30 (83)	18 (20)	<0.001
tment initiation at 3 mo	14 (38.9)	12 (13.5)	0.003
tment initiation at 6 mo	17 (47.2)	17 (19.1)	0.003
nition of abbreviation: AFB = ac es indicate n (%). ner's exact test (two-sided).	d fast bacilli.		



### Accreditation

Final Report: August 2021 — August 2022



National Jewish Health is accredited with Commendation by the Accreditation Council for Continuing Medical Education (ACCME) to provide continuing medical education for physicians. The NJH Office of Professional Education produced and accredited this program and adhered to the updated ACCME guidelines.

#### **Live Webinars**

National Jewish Health designates each live activity for a maximum of 0.5 *AMA PRA Category 1 Credit*™. Physicians should claim only the credit commensurate with the extent of their participation in the activity.

#### **Live Twitter Chats**

National Jewish Health designates each Other activity (social media discussion) for a maximum of 0.5 *AMA PRA Category 1*  $Credit^{TM}$ . Physicians should claim only the credit commensurate with the extent of their participation in the activity.

