

NTM Research Consortium Meeting Agenda

Friday, November 4th, 2016

Penner Ash Wine Cellars, Newberg OR

8:00am Shuttle from Inn at Red Hills to Meeting

8:30am – 9:00am

BREAKFAST: Continental Breakfast served in Meeting Room

9:00am – 9:15am

Welcome, Introductions and Meeting Objectives
Kevin Winthrop, OHSU

9:15am – 10:35am

NTM Research Consortium site updates (20 min each)
Clofazimine clinical trial, animal modeling, NW NTM Biobank/immunology
Kevin Winthrop and David Lewinsohn, OHSU

CF/NTM Research and Development Program
Charles Daley, National Jewish Health

Clinical cohort and isolate repository
David Griffith, University of Texas at Tyler

NIH biorepository, biomarkers, population-based studies
Kenneth Olivier, NHLBI

10:35am – 11:05am

Round table discussion: harmonization of repositories

BREAK

11:20am – 11:40am

Bronchiectasis and NTM Research Registry
Timothy Aksamit, Mayo Clinic

11:30am – 11:50am

Ontario laboratory-based studies and surveillance
Ted Marras, University of Toronto

11:50am – 12:20pm

Clinical trial design, TB drug candidates
Charles Daley, National Jewish Health

LUNCH: Catered buffet served in meeting room

1:30pm – 1:50pm

Patient reported outcomes: NTM Symptom Module
Alexandra Quittner, University of Miami

1:50pm – 2:10pm

Patient perspective
Amy Leitman, NTM Info & Research and local patients

2:10pm – 2:20pm

PCOR/NTM Research Roadmap update
Emily Henkle, OHSU

BREAK

2:30pm – 4:30pm

Round table discussion: future collaboration planning

WINERY TOUR 4:30pm / HORS D'OEUVRES 5:30pm/ DINNER 6:30pm-8:30pm

8:45pm

Shuttle back to Inn at Red Hills

2016 NTM Research Consortium Annual Meeting Attendees

Friday, Nov 4th - 8:30am-8:30pm

Penner Ash Wine Cellars, Newberg, OR

FIRST	LAST	Prof.Deg.	Title	INST/ORG
Timothy	Aksamit	MD	Pulmonologist; President of Bronchiectasis Research Registry	Mayo Clinic
Luiz	Bermudez	MD	Professor and Head, Department of Biomedical Sciences	Oregon State University
Sabina	Blizzard		Senior Research Assistant	Oregon Health & Science University
Joseph	Clock		Research Assistant	Oregon Health & Science University
Chuck	Daley	MD	Professor of Medicine; Chief, Div of Mycobacterial and Respiratory Infections	National Jewish Healthcare
Barbara	Fawcett		Bronchiectasis patient	
John	Fawcett		Bronchiectasis patient spouse	
John	Fowler	MD	Pulmonologist	The Vancouver Clinic
David	Griffith	MD	Professor of Medicine; Interim Director, School of Medicine Pulmonary Division	University of Texas Northeast
Emily	Henkle	PhD, MPH	Senior Research Associate	Oregon Health & Science University
Erin	Hernandez		Senior Research Assistant	Oregon Health & Science University
Amy	Leitman	JD	Director of Advocacy & Policy	NTM Info & Research
David	Lewinsohn	MD, PhD	Professor of Pulmonary and Critical Care Medicine	Oregon Health & Science University
* Francisco	Leyva	MD	Medical Officer, Div of Microbiology and Infectious Diseases	National Institute of Allergy and Infectious Diseases (NIAID) /NIH
* Dikoe	Makhene	MD	Medical Officer, Div of Microbiology and Infectious Diseases	National Institute of Allergy and Infectious Diseases (NIAID) /NIH
David	Mannino	MD	Professor of Medicine, Div of Pulmonology, Critical Care, and Sleep Medicine	University of Kentucky College of Public Health/COPD Foundation
Ted	Marras	MD	Respirologist; Director, Nontuberculous Mycobacterial Clinical and Research Program	University of Toronto/Toronto Western Hospital
Megan	Morgove	MS	Research Associate	Oregon Health & Science University
Kenneth	Olivier	MD, MPH	Senior Clinician, Pulmonary Clinical Medicine	National Institute of Allergy and Infectious Diseases (NIAID) /NIH
Wesley	Plinke		MPH Student, Study Coordinator	Oregon Health & Science University
* Becky	Prevots	PhD, MPH	Epidemiologist and Associate Scientist; Chief, Epidemiology Unit	National Institute of Allergy and Infectious Diseases (NIAID) /NIH
Alexandra	Quittner	PhD	Professor of Psychology, Pediatrics and Otolaryngology	University of Miami
Gregory	Ranches	MD	Pulmonologist	Oregon Health & Science University
Sam	Ruimy		Senior Research Assistant	Oregon Health & Science University
Sarah	Siegel	MPH	Research Associate	Oregon Health & Science University
Luke	Strnad	MD	Assistant Professor of Medicine Division of Infectious Diseases	Oregon Health & Science University
Kevin	Winthrop	MD, MPH	Professor of Public Health; Associate Professor of Infectious Diseases, Ophthalmology; Director,	Oregon Health & Science University

*attending via webinar

NTM Research Consortium Meeting Agenda

Friday, November 4th, 2016

Newberg OR

Attendees: see attached *list*

NTMRC Site Updates

OHSU (Kevin Winthrop/Emily Henkle/Dave Lewinsohn)

Oregon NTM Cohort

- Results presented for mortality/clinical outcomes; papers in submission

NW NTM Biobank

- 350 enrolled, funded through June 2017
 - 300 pulmonary NTM
 - Rest extrapulmonary, pulmonary controls, healthy controls
- Aims to establish cohort, pilot immunology (MAITs/cytokine profiles)
- Clinical data, blood, QOL-B/NTM module, GAD7/PHQ-8
- Utilizes REDCap database

PCORI Non-CF Bronchiectasis Comparative Effectiveness and Safety

- Non-CF bronchiectasis cohort (full Medicare, 2006-2014)
- Safety and effectiveness of inhaled corticosteroids, macrolides, inhaled antibiotics
- Primary outcomes hospitalized pneumonia, NTM infection
- Ongoing, beginning descriptive analysis

Animal model

- Successful infection of 3 macaques

Pilot immunology

- Focus on mucosal-associated invariant T (MAIT) cells
- Peripheral MAIT cells missing in those with TB
- Pilot results: MAITs reduced in males and individuals with COPD

National Jewish (Chuck Daley)

Adult day unit

- Now incorporating QOL-B/NTM module and AE monitoring via Redcap electronic entry tool
- Looking at gene expression in sputum, capture blood at baseline/6m to look for biomarkers of injury/assoc. with AEs optic neuritis
- 500 patients with specimens/NTM with electronic medical record data

Colorado Cystic Fibrosis Research and Development Program

- Three research cores
 - NTM Culture, Biorepository, and Coordinating Core
 - Clinical Research Core
 - Molecular Core
- Pilot innovation funding: free whole genome sequencing on all isolates from CF patients
 - 393 isolates collected
 - Goal to identify patient-to-patient transmission
 - Working on linking samples to CF registry (complicated)
- PREDICT: Pre-treatment natural history protocol, quarterly visits after isolating NTM
- PATIENCE: Optimal treatment
- Planning to expand PREDICT/PATIENCE to other sites

Biorepository

- Enrolling patients in genetic biobank
- Save samples of any sample collected for clinical use (blood, urine, tissue, sputum, etc.)

NIH (Ken Olivier)

NHLBI clinic

- Most patients enrolled on a protocol- types of data collected are standard, though the frequency is not (sicker patients have more follow up)
- Most relevant protocols
 - Idiopathic bronchiectasis (stratified by sex, NTM)
 - CF registry started in 2010
 - Increasing NTM capture

NIAID Epi unit (*Becky Prevots*)

- Access to large national datasets
 - 5% Medicare sample, AHRQ utilization,
 - CF Foundation Registry
 - Hospital, HMO databases
 - Linkage to other Federal databases: NOAA, FAA, USGS
- Recent study
 - Hawaii- epi of NTM in state with highest rate of NTM.
 - Kaiser EHR data
 - Explore host/environmental factors

UT (David Griffith)

Clinical cohort

- Excel database with 1000 patients going back to 1970s.
- Biorepository: Save one isolate per patient (all patients)

Animal models

- Marmosets, pig (CF)

Invited Presentations

Bronchiectasis and NTM Research Registry (Tim Aksamit)

- Publication describing baseline data accepted at Chest
- Moving forward:
 - PPRN
 - Linkage to CMS, electronic health records

Ontario surveillance (Ted Marras)

- Laboratory based studies-
- Recent studies
 - Obstructive lung disease and NTM risk
 - Population-based, matched cohort study of survival after pulmonary NTM (by species, case or isolate)

Clinical Trials- lessons from TB experience (Chuck Daley)

- Timeline is long for new drugs (keep working on these)
- In interim, work on improving regimens
- Adaptive design allows for efficient study
- Think about pragmatic trials

NTM Module development (Alexandra Quittner)

- NTM Module developed according to FDA guidance on PROs (2009)
- Preliminary psychometric analyses complete
- Next steps: completing final psychometric analysis, determining MID

NTM Patient-centered research priorities from November 2015 meeting (Emily Henkle)

- Workshop report published: Ann Am Thorac Soc. 2016 Sep;13(9):S379-84.
- Summarized 13 patient-centered priorities
 - Working on person-to-person transmission, NTM symptom module, anxiety/depression impacts, correlation of HRQoL and clinical outcomes
 - Several still need to be addressed
 - Role of GERD
 - Validate molecular diagnostics
 - Comparative effectiveness of exercises/lung clearance devices
 - Understand who needs/benefits from therapy
 - New antibiotics, shorter courses of therapy duration
 - Disease severity index
 - Biomarkers

Discussion: Next steps for consortium

- No approved treatment for NTM
- Need for large, pragmatic trials balancing concerns for capacity
 - Pursue adaptive design, comparing regimens
- Need to organize sites- cultures, follow-up
 - Identify quality labs for diagnosis/culture
 - Agree on initial evaluation/standard of care at “centers of excellence”? Work with NTMir?
 - Then expand to additional sites/physicians
- Common measures (include above standard of care, plus additional relevant to specific research questions)
- Explore common data models
 - PCORnet, Mini-sentinal, HMORN, etc
- Need for biomarkers
- NIH grants feasible, need to be hypothesis driven
- Models for consortium formation/growth
 - PCD rare disease network
 - CF Clinical Centers networks/Registry