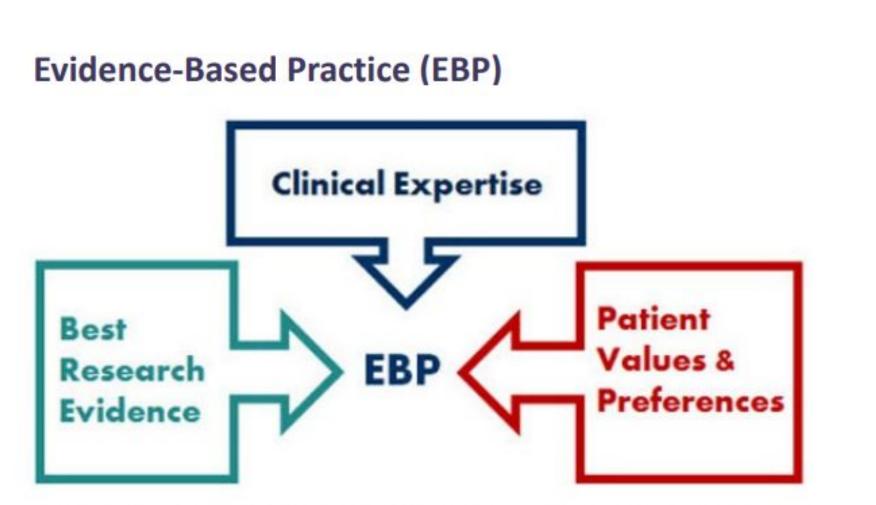
In the light of the Case series published-Nocardia infection causing non-healing surgical wounds: A case series from Bangladesh

Prof. Dr. Afzalunnessa Binte Lutfor Head, Microbiology Department Ad-Din Women's Medical College, Moghbazar Former Research Associate Tohoku University, Japan Former Professor and Head of Dept. Microbiology, SSMC and SZMC

Evidence Based Practice is the idea that occupational practices ought to be based on scientific research

Criteria of for EBM or EBP

Clinical expertise/expert opinion Evidence (external or internal) Client/patient or/caregiver perspective https://www.asha.org>research>ebp Father of early modern medicine--Ibne Sina Father of evidence based medicine- Archie Cochrane



"Evidence based medicine is the conscientious, explicit, and judicious use of current best evidence in making decisions about the care of individual patients. The practice of evidence based medicine means integrating individual clinical expertise with the best available external clinical evidence from systematic research." Sackett et al., 1996

http://guides.mclibrary.duke.edu/ebmtutorial/home https://www.bmj.com/content/312/7023/7



Louis Pasteur (1822-1895)-Father of Microbiology

In the field of observation, chance favours only the prepared mind. Louis Pasteur

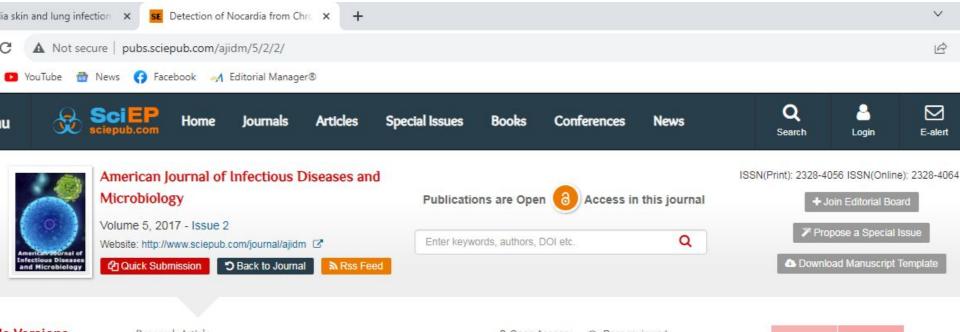
More science quotes at Today in Science History todayinsci.com

The Journey of Nocardia detection started in 2016 – A Vision from a great surgeon Late Prof. A. A. Ashraf Ali and the Mission taken by the Microbiology Department



A Nocardiosis case- Which door to knock?

Published in 2017





RIS

Both diagnostic services and clinical practice depends on observation at the beginning



Dependency of the Laboratory Division and our expectations Depends on Our Expectations

- Doctors who caters direct service to patients
- Patients who voluntarily collect and supply
- Receptionist/Phlebotomis t who receives or collects sample

- Proper request
- Additional request
- Correct samples
- Authentic samples so that does not fall into the "sample rejection" criteria
- i.e., swab/pus/discharge not tissue in formalin

Background

- Nocardia can be introduced accidentally or by unhygienic habit leading to non-healing surgical wounds. It should not be an endogenous source.
- This is enigmatic for clinicians and troublesome for patients because of festering infections for months or years as culture needs >5 days or extended incubation and treatment needs >6 months antibiotic- both of which exceptional.
- With timely diagnosis and by appropriate treatment, *Nocardiosis can be cured*.
- Delayed or missed diagnosis can lead to the development of antimicrobial resistance.

Objectives

Eliminate diagnostic dilemma

Disseminate to the doctors

Treatment before unwanted delay

Methodology

- All patients were referred to the Microbiology Laboratory of AWMCH and different hospitals or labs of Dhaka and other districts.
- Detailed history with cell phone noted.
- Antibiotic withdrawn 3-7 days before sample collection
- Samples from wounds were collected in the laboratory under supervision.
- *Nocardia* identification and susceptibility testing were carried out by standard procedure with a little twist or in house adaptation for the extended culture.

Special measures taken in diagnosing Nocardia spp.

- Referred selective cases
- History suggestive
- Antibiotic discontinued
- Sample taken at the lab
- Multiple sample and multiple staining, culture
- Extended culture with special care to prevent drying and contamination
- Call to collect report as may take 7-10 days
- Feedback from the patient regarding treatment and recovery through phone or app to collect pictures.

Case summary:

Mrs. X, 26 year old, normotensive, a graduate from Uttara presented with the complaints of -

• Chronic ulcer on abdominal scar for 11 months.

She stated that she had elective LUCS in 2020. After 15 days since the surgery she developed infection on wound site and was given antibiotic according to wound culture. But despite repeated courses of antibiotics for 6 months, wound was not healing. Then six months back, she was diagnosed with skin TB and sinus formation in the wound site. Sinectomy was done and was given modified CAT-1 regimen with moxifloxacin. As the wound still persists, she was referred to pulmonologist for further management.

She had a H/O incomplete abortion in 2019 due to maternal Rubella and CMV infection and had developed gestational diabetes and Covid-19 during pregnancy.

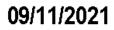
On general examination: Pulse 90/min, BP 90/60 mm of Hg, Temp – normal, $SaO_2 - 97\%$ without Oxygen, six min walk test: No desaturation, weight 65 kg, Respiratory system examination: normal

Wound: multiple non healing ulcer, irregular margin, with discharge from some lesions

Family history:	NAD
Personal history:	Nothing significant
Socioeconomic history:	middle class
Occupation:	Private employee (desk job)
Travel H/O:	NAD
Exposure H/O:	NAD
Previous illness:	Maternal Rubella, HSV-1,Rubella infection in 2019, Covid-19 in
	2020, gestational diabetes
H.O allergy:	NAD
Smoking History	Non-smoker
ongoing Treatment	
Previous treatment	Extended CAT-1 regimen for 9 months, Clindamicin,
	flucloxacillin

736 Mob. : 01637-216322



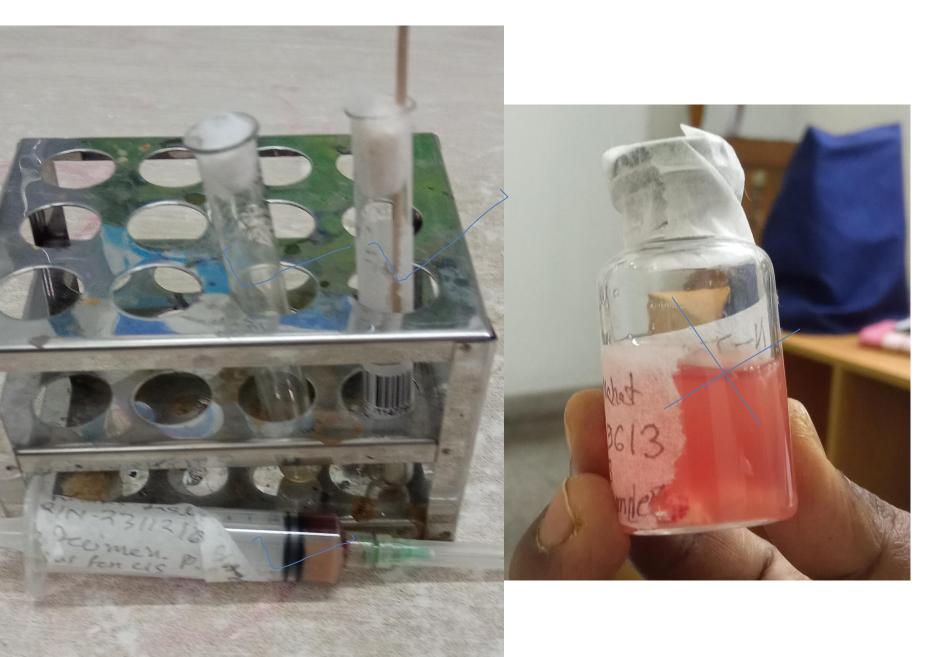


Diagnosis	
1. Post LUCS chronic discharging sinus	
Investigation	
1. Multiplex PCR of TB & NTM (BSMMU)	
2. Culture for Nocardia from biopsy sample (Prof. Afzalunnessa, Ad Din Medical	
College)	
Treatment	

Diagnosis- based on 3 criteria

Gram stain MZN stain ZN stain Culture 1.Growth positive plus
2. One Gram stain and
3. One MZN stain must have positive findings

Unavoidable-Sample rejection



Acknowledgement: Technicians, & doctors for kindly collecting samples and detailed history taking





Sample collection-points to ponder

Beware of super added infected condition



Discontinue before collection Antibiotic for 4 days and Anti TB drugs for 7 days- Ref.: Afzalunnessa et al., *American Journal of Infectious Diseases and Microbiology*, 2017, Vol. 5, No. 2, 80-86



Methodology

- Detailed History
- Histopathology /PCR report
- Sample
- Gram stain
- ZN stain
- MZN stain
- Culture
- Subculture
- Sensitivity
- Storage



Prevent drying and contamination



Gram positive filaments in cocco-bacillary form



Findings on MZN stain of primary culture, like this Makes your day



Finding this on MZN stain



Microscopy-a tedious relentless search

- Gram staining
- Ziehl-Neelsen staining
- Modified ZN staining-3%
 Sulphuric acid





Variations in growth of Nocardia



rowth of Nocardia spp. on Blood Agar plate



a Binte Lutfor et al. Detection of Nocardia from Chronic Skin and Lung Infections in E purnal of Infectious Diseases and Microbiology, 2017, Vol. 5, No. 2, 80-86. doi:10.126

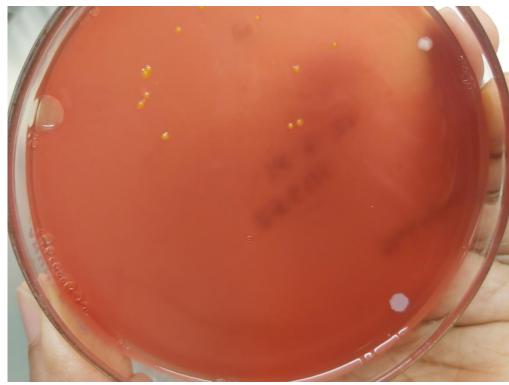
r(s) 2015. Published by Science and Education Publishing.



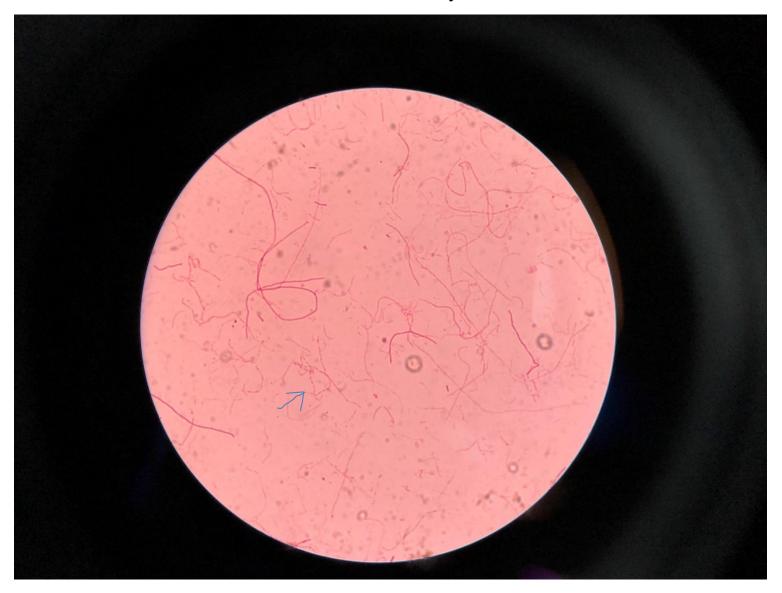
Pick up the right colony from mixed growth



Suspected colony grown after 3 days incubation



Acid fast bacilli in cocco-bacillary form



Patient 11

A woman aged 56 years came with discharging as well as healed sinuses on the <u>abdominal wall</u> (image 8). She underwent a <u>hysterectomy</u> and after the operation, her wound never healed. For one and a half years, she was treated with multiple antibiotics, antifungal and even anti-tubercular drugs. *Nocardia* spp. was isolated from the sinus above the stitch area and the patient was cured with co-trimoxazole taken for one year.



Download : Download high-res image (156KB) Download : Download full-size image

Patient 7

A 60-year-old male presented with a few discharging sinuses following exploratory surgery on a nodule in the right infra-mammary region (image 5). Culture of pus from the discharging sinus was identified as *Nocardia* spp. Doxycycline was given according to susceptibility pattern for 12 months, and the patient was cured.



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Figure 5. Few discharging sinuses which developed following exploratory surgery on a nodule in the right inframammary region of a 60-year-old businessman. A case of RTA, taken antibiotic, antifungal, anti-TB drugs but after diagnosis of Nocardiosis

cured with doxycycline taken for a year





Afzalunnessa et al., International Journal of Infectious Diseases 110 (2021) 272-278. https://doi.org/10.1016/j.ijid.2021.07.028







Figure 11. A 35-year-old businessman presented with a swelling in the left submandibular region with multiple sinuses around the swelling.



Figure 12. Multiple discharging sinuses are visible around the appendectomy incision of a 36-year-old female.

treatment for 3 months without much improvement. Burkholderia pseudomallei was identified from the pus at the base of the swelling and culture from the discharging sinus near the chin found Nocardia spp. He was given 14 days of ceftriaxone for Burkholderia pseudomallei and 6 months of doxycycline for Nocardia spp. The patient noticed remarkable improvement after six months and is still under antimicrobial therapy.

The only immunocompromised patient- a child with X-linked agamma-globulinemia





More cases



Figure 3. A discharging sinus on the right side of neck in a 32-year-old male banker is visible along with scars of some healed sinuses.



Figure 4. Anterior abdominal wall of a 47-year-old female shows a non-healing wound above the scar of herniotomy and hernioplasty operation.



Figure 6. Lower abdominal wall of a 30-year-old female showing a discharging sinus above the horizontal incision for lower uterine caesarian section.



Figure 7. Lower abdominal wall of a 34-year-old female showing a discharging sinus above the horizontal incision for lower uterine caesarian section. Some suture material is still visible.



Development of infection after surgical procedure-a thought provoking issue for the Microbiologist-IPC team?

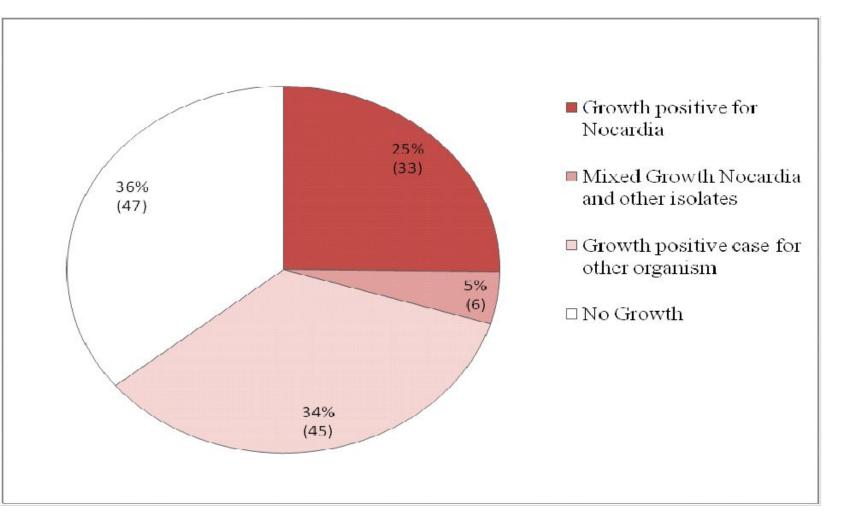
H/O laparoscopic surgery with port infection



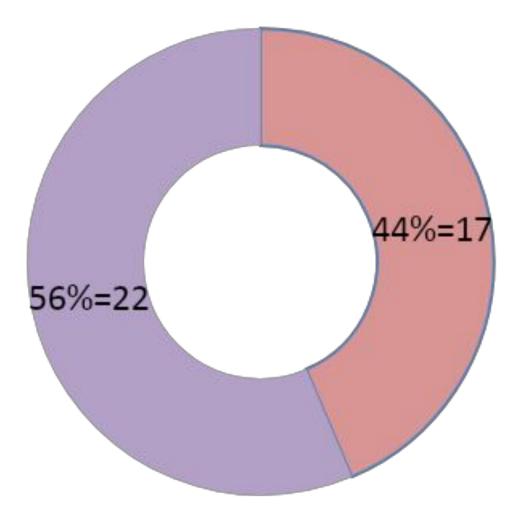
H/O surgery with emerging sinuses and healed marks around



Results (2016 to 2020) Among total referred cases (131), 39 (33+6) *Nocardia* spp.

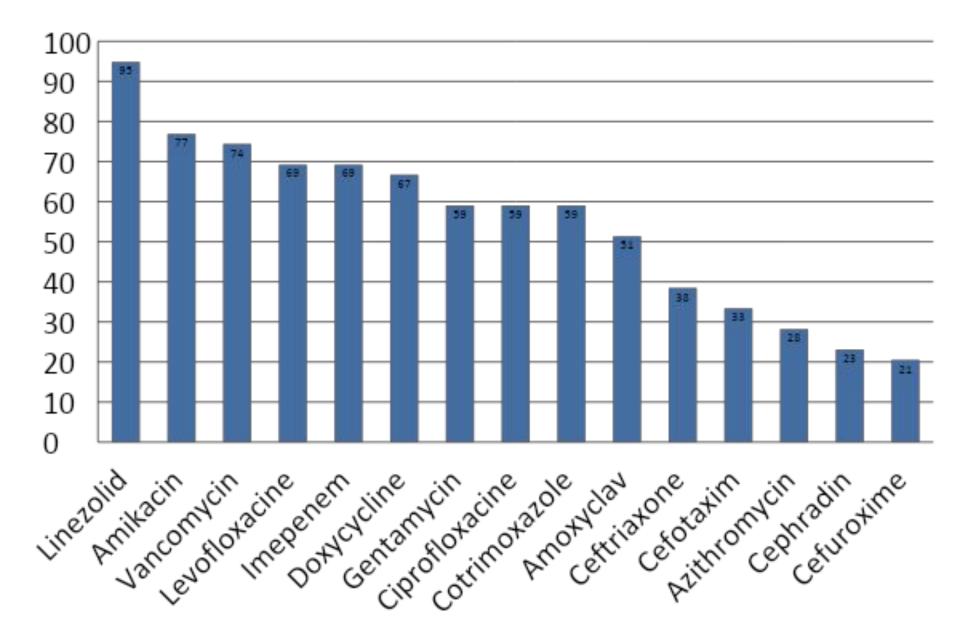


Among total 39 Nocardiosis cases, 17 (44%) have been prescribed or taken anti-TB drugs



Suspected TB or having antitubercular drug = 17 Not having antitubercular drug = 22

Sensitivity pattern of Nocardia Spp.



Identifying *Nocardia* spp. -challenging but not impossible

Nothing is impossible, the word itself says **'I'm possible'!**

– Audrey Hepburn

AZQUOTES

Conclusions

• Non healing wounds following surgery or trauma should be explored for Nocardiosis

 Granulomatous lesion should also be ruled out for Nocardia

 Resistance to commonly used antimicrobials is on the rise

Take home message

- 1. Prevent the possible breach of sterilization protocol of laproscopic and surgical instruments.
- a) GTA (Guteraldehyde) 2% for 30 mins isn't sufficient, P^H (3-4) needs to be assessed everyday (half life 20-28 days) or at least 8 hrs soaking is needed.
- b) OPA (Orthophthaldehyde) or Cidex for 30 mins.
- 2. Use of 10% house bleach for floor/ surfaces(Na-hypochloride
- 3. Use of autoclavable/disposable instrument.
- 4. Plasma sterilizer or Ethylene Oxide in between consecutive surgery or procedure.
- 5. Enzyme or Ultrasonic technology in device cleaning with autoclaved water.

Reported by `Umar (RA) said: I heard Messenger of Allah (peace be upon him) saying: "If you all depend on Allah with due reliance, He would certainly give you provision as He gives it to birds who go forth hungry in the morning and return with full belly at dusk". [Tirmidhi] Sometimes, we think it is our careers that put food on our table, let us remember that all provisions come from Allah. Say Alhamdulillah

