



Melioidosis

“A Neglected Topical Disease”

Sotharith BORY, MD., MSc.

Internist-Infectious Diseases Physician

Melioidosis?



TB?

Cancer?



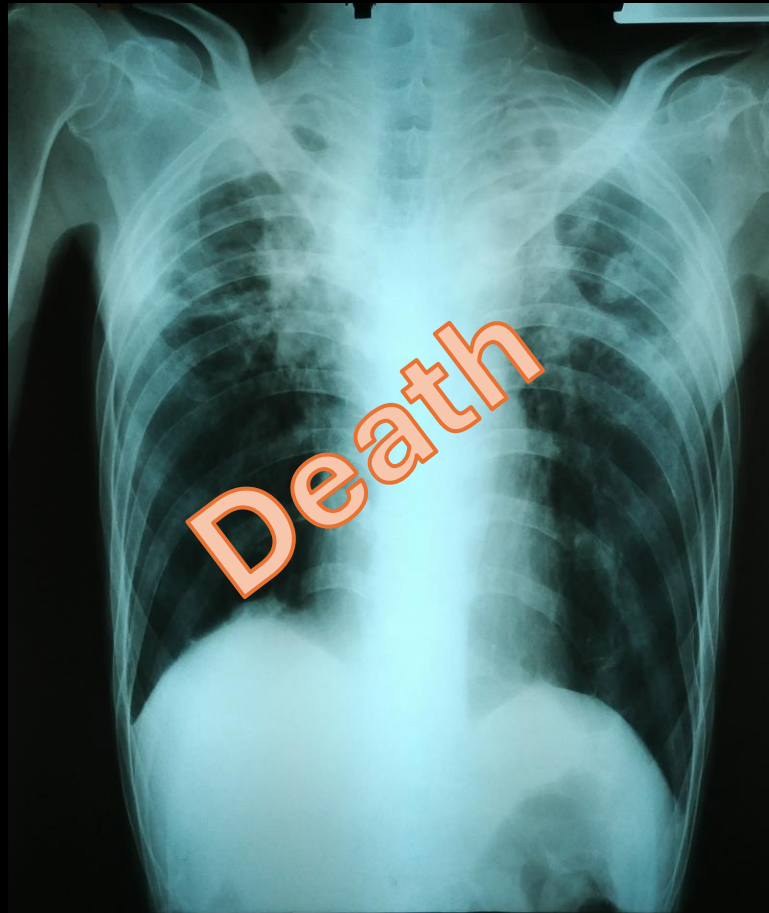
Case 1: 58-year-old male rice farmer, diagnosed with “TB” on chest X-ray in provincial hospital. Wanted a second opinion – self referred to Phnom Penh.

* 1-month history of fever, cough, chest pain, difficulty in breathing, headache, and weight loss.

* AFB: negative

* Sputum culture and X-pert: negative

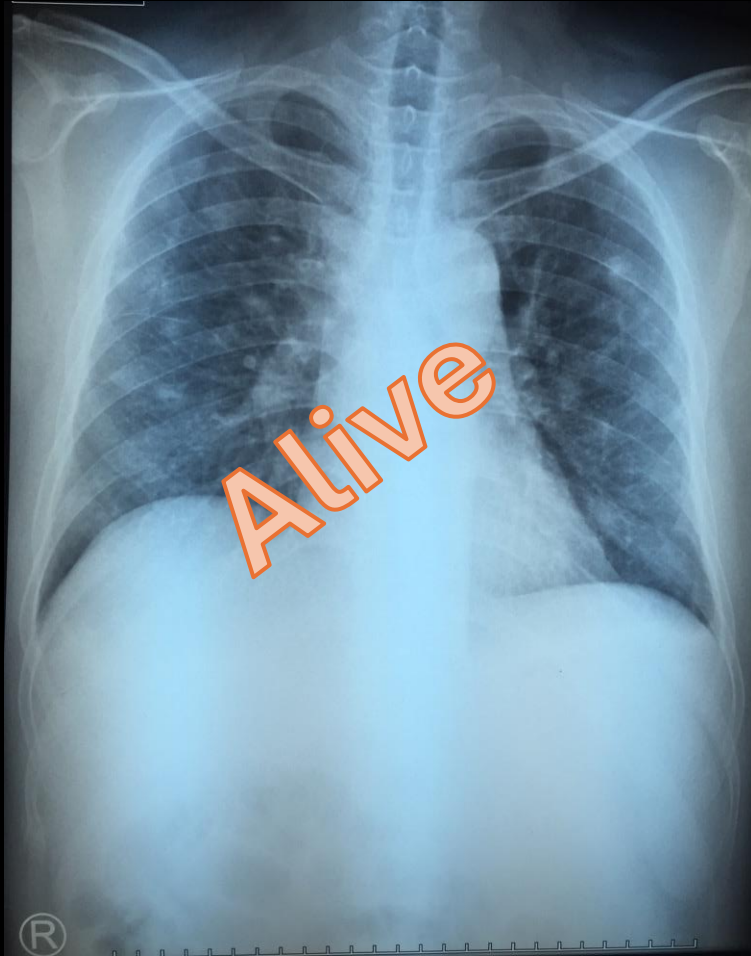
➔ **Blood culture: *Burkholderia pseudomallei***



Case 2: 86 yo M, Ex-farmer from Kampong Chnang

- 1 year history –fever with chills, cough with sputum, dyspnea
- > 6 months earlier, AFB negative, but CXR suggestive of TB → Treated for TB (RHZE) 6 months but no clinical improvement. Condition worsened (worsening dyspnea, cough with yellow sputum) → Calmette H

→ Sputum and Blood culture: **B. pseudomallei**

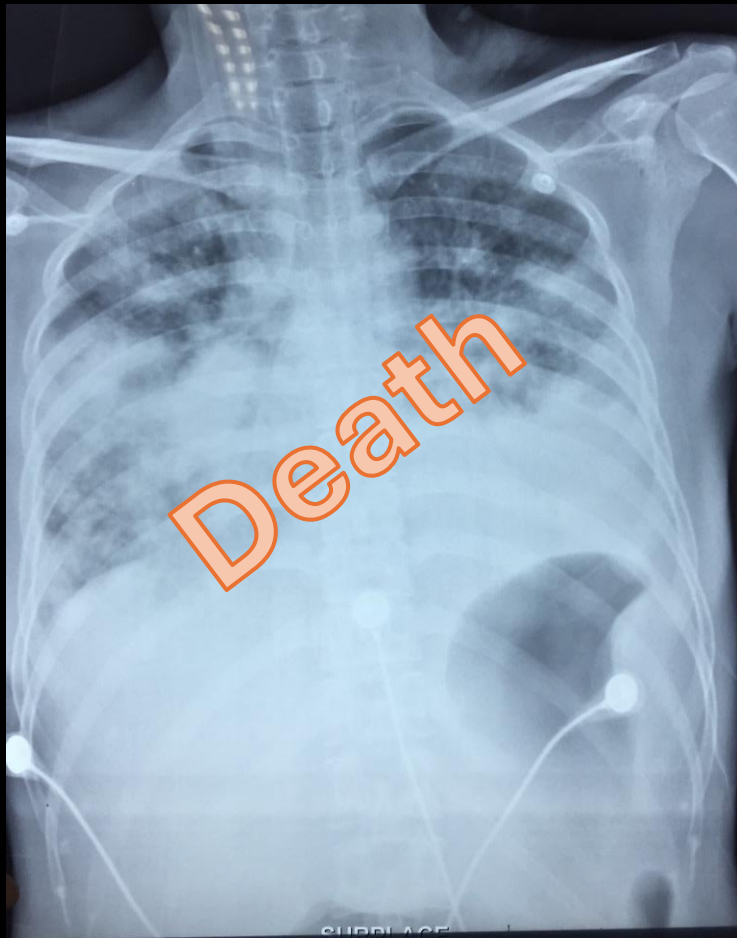


Case 3: 48 year old man, from Prey Veng province, prolonged fever with productive cough → Treated with anti-TB drugs (RHZE) - finished 1 month before admitted to our hospital.

* Diabetic

- Sputum: AFB Negative
- Bacterial Culture: Negative
- Gene X-pert: Negative

→ **Blood culture: *B.pseudomallei***



Case 4: 41 Year old female, hair-dresser from Thboun Khmom province,

* Presented to Calmette hospital with prolonged fever and chest-pain.

* Pulmonary tuberculosis:

- Treated July 2017 (RHZE- 6 mths) Smear (U/K)

- Treated March 2018 (8 mths), Smear (-)

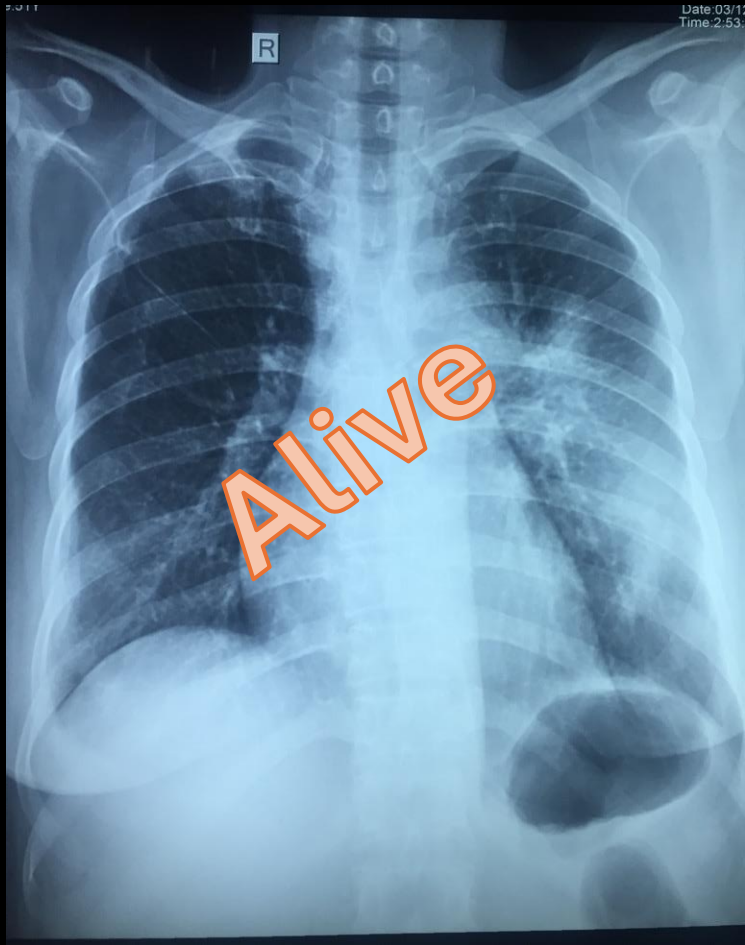
- Diabetes: Unknown

* **Admitted with suspect of Bronchial tumor (non smoker)**

- Sputum: AFB: (-), Gene X pert: (-).

- Culture (Bacterio): Negative

➔ **Blood culture: *B. pseudomallei***



Case 5: 55 Year old female teacher from Svay Reang province,

* Presented with fever and cough more than 2 months with weight loss.

- AFB Smear: negative

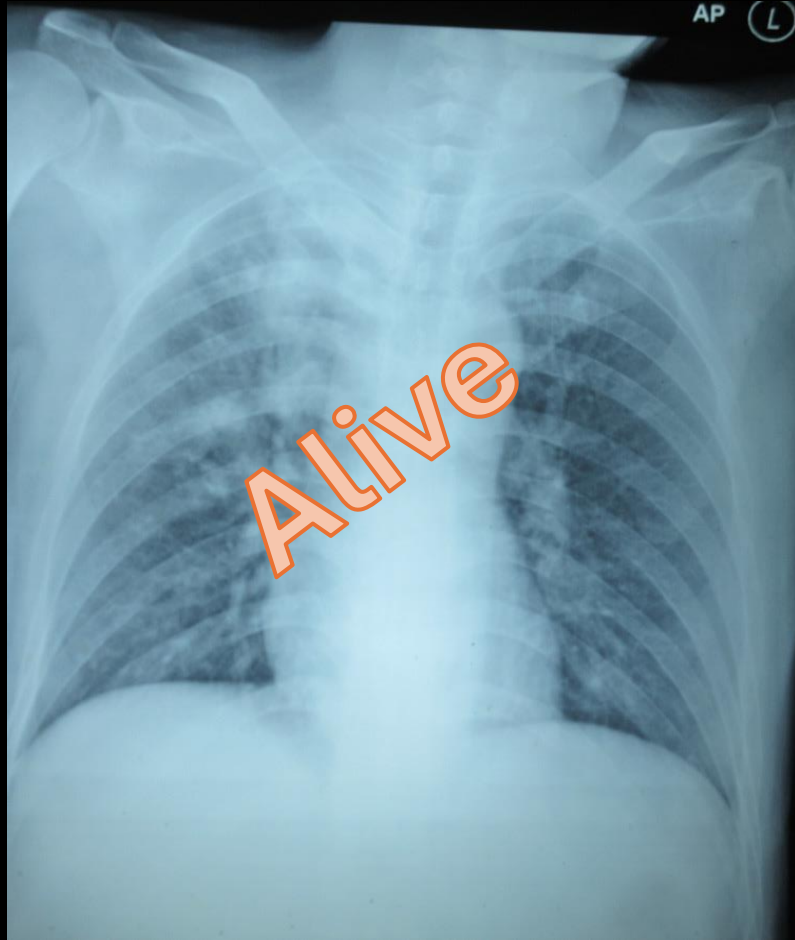
➔ **Sputum & Blood culture:**
Burkholderia pseudomallei



Case 6: 41 Year old male
carpenter from Pursat province,
* Presented to Calmette
hospital with prolonged fever
and dyspnea

* Past medical history: Diabetes
* Pleural fluid: negative
(culture and X-pert)

➔ **Blood culture: *Burkholderia pseudomallei***



Case 7: 39 Year old male, motor taxi from Kompot province.

- * Presented to Calmette hospital with prolonged fever and cough (more than 1 month) and weight loss

- * Smear for AFB and Gene Xpert from sputum: negative

- * Sputum culture: not done

→ **Blood culture: *B. pseudomallei***



Case 8: a 49-year-old female presented with a productive cough, fevers and weight loss.

- * More than 10 years ago - treated twice for pulmonary TB “national TB protocol” → improved a little only and went home, but symptoms persisted
- * Sputum AFB: (-)
- * Gen X-pert: (+)

→ **Blood culture, sputum culture: *B. pseudomallei***

Tuberculosis mimicked by melioidosis

K. Vidyalakshmi,* M. Chakrapani,† B. Shrikala,* S. Damodar,† S. Lipika,* S. Vishal†

*Department of Microbiology, †Department of Medicine, Kasturba Medical College, Mangalore, India

Melioidosis should be ruled out in patients with suspected TB while instituting presumptive anti-tuberculosis treatment in areas where both diseases are prevalent.

Fever in a diabetic patient with high ESR and neutrophilic leucocytosis should raise suspicion of melioidosis, even if radiography or cytopathology is indicative of TB. The suspicion of *B. pseudomallei* should especially be raised when *Pseudomonas*-like isolates are resistant to aminoglycosides and/or colistin.

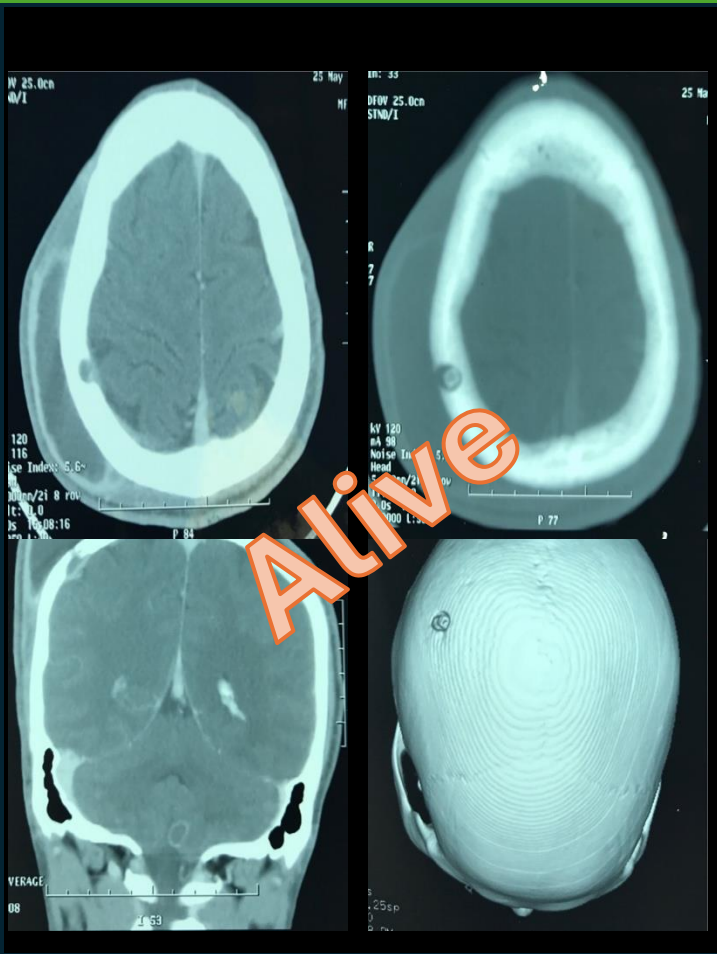
RESEARCH ARTICLE

Open Access

Pulmonary melioidosis in Cambodia: A prospective study

Blandine Rammaert¹, Julien Beauté¹, Laurence Borand¹, Sopheak Hem¹, Philippe Buchy¹, Sophie Goyet¹, Rob Overtoom², Cécile Angebault¹, Vantha Te³, Patrich Lorn Try⁴, Charles Mayaud⁵, Sirenda Vong^{1*} and Bertrand Guillard¹

We suggest, therefore, that **pulmonary melioidosis should be considered in every patient with TB-like chest X-rays and negative AFB smears in Cambodia.**



42 male, farmer from Kompong Chnaing province, **Scalp swelling**, tenderness with fever for about **4 months**.

Past medical history:

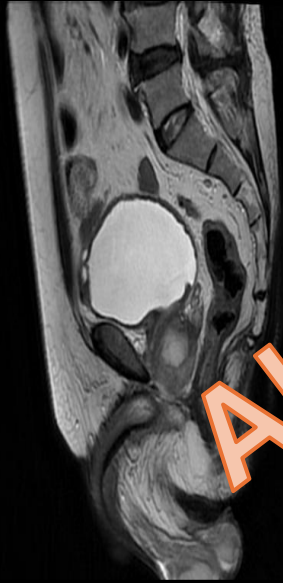
- Uncontrolled diabetes for 2 years
- No past history of head trauma

An operation was done on **day 2**

- Pus collection under the scalp
- The bone was affected → culture



→ **Both blood and pus culture (+) for B.P**



Alive



75 male, rice farmer from Kompong cham province, came to Calmette Hospital for fever with rigors and dysuria for 1 month

Past medical history:

- Steroids using for the problem of joint pain
- No profile of diabetes before hospitalization

Result:

- **Urine culture, Pus culture (+) B.P**
- **Blood culture also (+) for B.P**

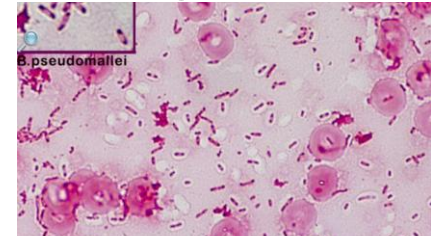
MELIOIDOSE

What is Melioidosis?

- ❖ Infectious disease caused by the Gram-negative bacterium *B. pseudomallei*.
- ❖ It is found in soil and water, particularly in tropical and subtropical regions.

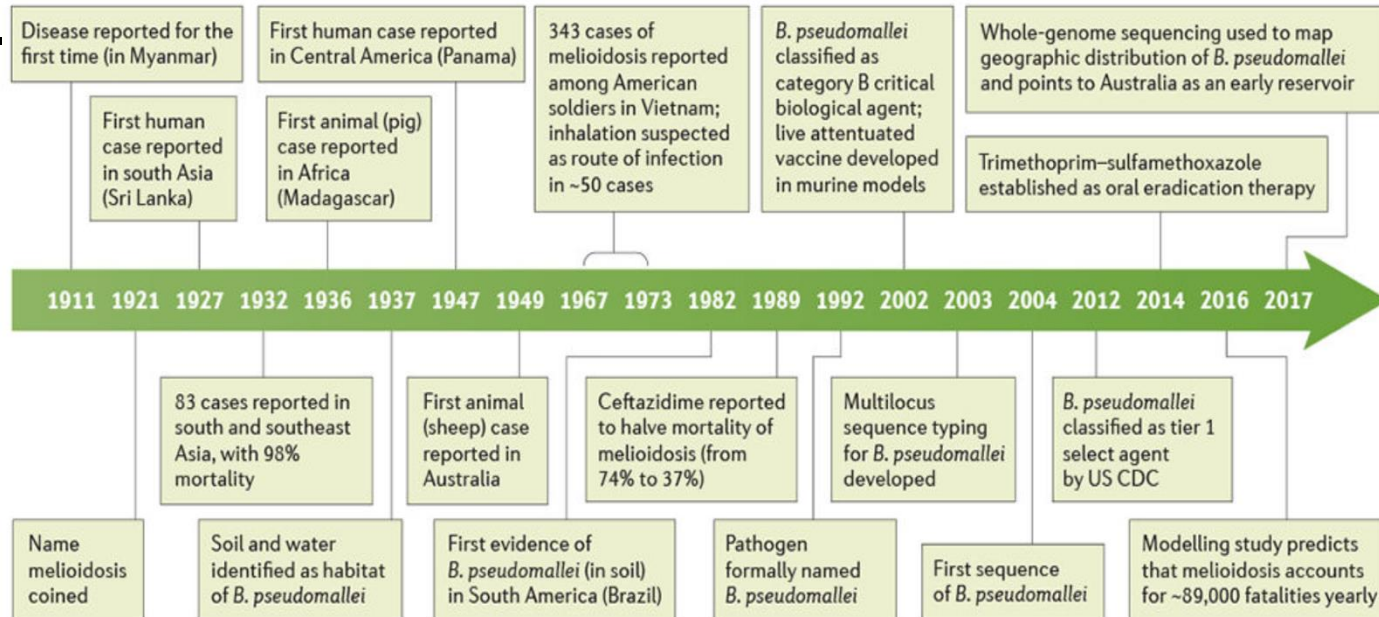
Transmission:

- ❖ Percutaneous inoculation (e.g., through skin abrasions)
- ❖ Inhalation
- ❖ Ingestion



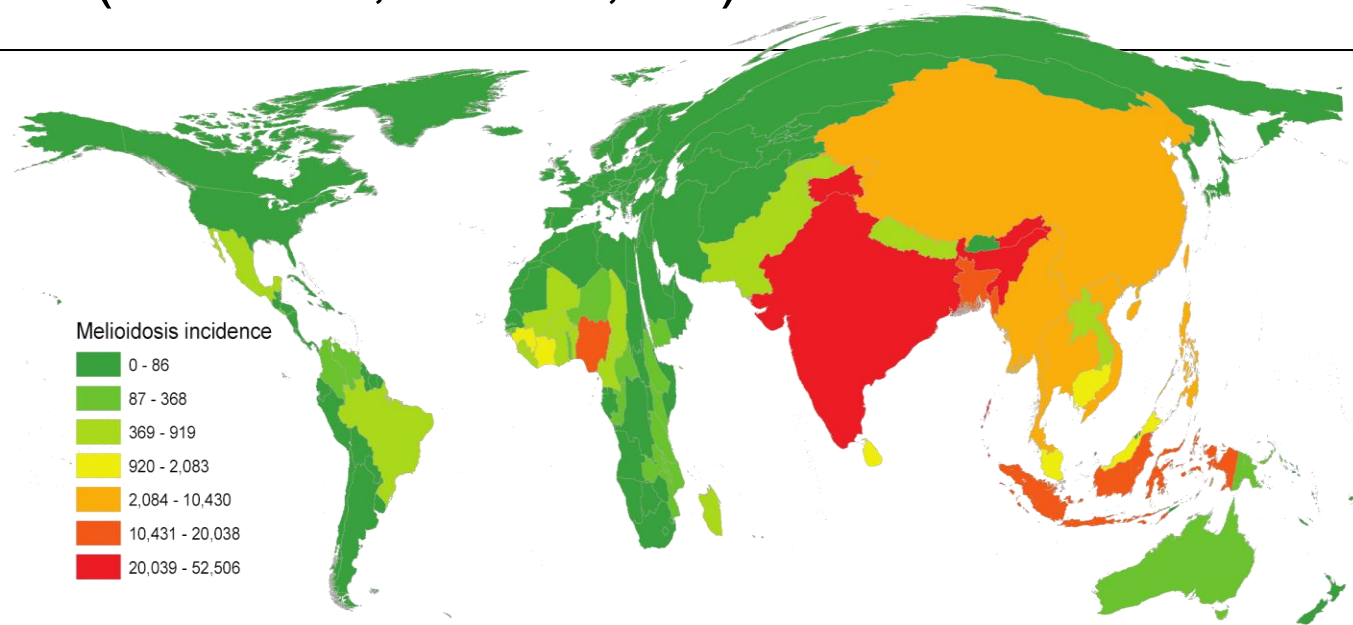
Back to the history

- ❖ First described in **1911** by **Whitemore** and his assistance **Krishnaswami** in **Burma** (now known as Myanmar).
- ❖ The name melioidosis was later coined by **A.T. Stanton** and **W. Fletcher** in **1921**.



Predicted global distribution of *Burkholderia pseudomallei* and burden of melioidosis (Nature Microbiology, 2016)

Main Result: We estimate there to be **165,000** (95%CI 68,000-412,000) human melioidosis cases per year worldwide, of which **89,000** (95%CI 36,000-227,000) die.



Cartogram of the incidence as a proportion of national geographical area in 2015; **44% is in South Asia**

Predicted global distribution of *Burkholderia pseudomallei* and burden of melioidosis


Direk Limmathurotsakul^{1,2,3}, Nick Golding¹, David AB Dance^{4,5}, Jane P Messina⁶, David M Pigott¹, Catherine L Moyes¹, Dionne B Rolim⁷, Eric Bertherat⁸, Nicholas PJ Day^{2,5}, Sharon J Peacock^{2,9,10}, and Simon I Hay^{1,11,12}

Disease	Predicted incidence	Predicted mortality	Predicted case fatality rate	Source of data
Tuberculosis	8,600,000	1,300,000	15%	WHO website
Malaria	219,000,000	660,000	0.3%	WHO website
2009 Pandemic Influenza A H1N1	N/A	284,000	N/A	LID 2012, 12(9)687-695
Melioidosis	165,000	89,000	54%	Nature Microbiology 2016
Leptospirosis	1,000,000	59,000	6%	PLoS NTD 2015
Severe Dengue	500,000	12,500	2.5%	WHO website

Countries (selected)	Predicted Incidence	Predicted Mortality	Predicted Mortality Rate
India	52,506	31,245	60%
Indonesia	20,038	10,224	51%
..
Thailand	7,572	2,838	38%
Myanmar	6,247	3,687	59%
Cambodia	2,038	1,149	56%
Lao PDR	420	260	62%

MELIOIDOSIS IN CAMBODIA

- First documented case **May 1928** – Russian residing in Bangkok visited Cambodia
- **1960 - 2 cases in animals** - Takeo and Pursat
- No further human reported cases **until 2005** – Angkor Hospital for Children, Siem Reap
- **In 2006**, Calmette Hospital diagnosed this diseases on the diabetes patients and this issue was presented as a talk In the event of Cambodge Santé 2008.



Mélioidose

(A propose de 6 cas de la mélioidose recueilli dans le service de Med. A , hôpital CALMETTE)

Dr.Ouk Vara, Touch Socheat*, Sin Bochevan*, Niv Rathvirak*, Hy Chanseila*, Keo Vannarith*, Dr.Ong Sopheavy, Dr.Ear Rattanak, Dr.Ros Sina, Dr. Lim Vadhana, Pr.Nhem Sophoeun, Pr.You Kim Yean et équipe de Med. A.

430

BULLETIN DE LA SOCIÉTÉ DE PATHOLOGIE EXOTIQUE

Communications

Un cas de mélioidose observé à Phnôm-Penh,

Par A. GAMBIER.

Dans la matinée du 9 mai 1928 la police conduisait à l'hôpital de Phnôm-Penh M. ALM... sujet russe habitant habituellement Bangkok, que son état de surexcitation faisait prendre pour un dément. A son entrée le malade avait une température de 40°1. L'examen somatique est absolument négatif. Pas d'hypertrophie de la rate et du foie. Pas de troubles intestinaux. Pas de réaction ganglionnaire.

Emergence of Pediatric Melioidosis in Siem Reap, Cambodia

Yos Pagnarith, Varun Kumar, Janjira Thaipadungpanit, Vanaporn Wuthiekanun, Premjit Amornchai,
Lina Sin, Nicholas P. Day, and Sharon J. Peacock*

Angkor Hospital for Children, Siem Reap, Cambodia; Mahidol-Oxford Tropical Medicine Research Unit, Faculty of Tropical Medicine, Mahidol University, Bangkok, Thailand; Department of Microbiology and Immunology, Faculty of Tropical Medicine, Mahidol University, Bangkok, Thailand; Center for Clinical Vaccinology and Tropical Medicine, Nuffield Department of Clinical Medicine, University of Oxford, Churchill Hospital, Oxford, United Kingdom; Department of Medicine, University of Cambridge, Addenbrooke's Hospital, Cambridge, United Kingdom

Burkholderia pseudomallei Antibodies in Children, Cambodia

Vanaporn Wuthiekanun,* Ngoun Pheaktra,†
Hor Putchhat,† Lina Sin,† Bun Sen,†
Varun Kumar,† Sayan Langla,*
Sharon J. Peacock,*† and Nicholas P. Day*†

F1000Research

F1000Research 2015, 3:302 Last updated: 15 MAY 2015

in
w
vic
oic

RESEARCH NOTE

REVISED Detection of *Burkholderia pseudomallei* in Sputum using Selective Enrichment Broth and Ashdown's Medium at Kampong Cham Provincial Hospital, Cambodia [v2; ref status: indexed, http://f1000r.es/5e9]

Somary Nhem¹, Joanne Letchford², Chea Meas¹, Sovannndeth Thann¹,
James C. McLaughlin², Ellen Jo Baron³, T. Eoin West⁴



CrossMark
click for updates



ELSEVIER

available at www.sciencedirect.com

ScienceDirect

journal homepage: www.elsevierhealth.com/journals/trst



A first report of pulmonary melioidosis in Cambodia

Rob Overtoom^{a,b,*}, Virak Khieu^b, Sopheak Hem^b, Philippe Cavailler^b,
Vantha Te^c, Sarin Chan^b, Phea Lau^c, Bertrand Guillard^b, Sirenda Vong^b

^aSwiss Red Cross, Takeo, Cambodia

^bInstitut Pasteur in Cam

Rammaert et al. BMC Infectious Diseases 2011, 11:126

http://www.biomedcentral.com/1471-2334/11/126



Infectious Diseases

RESEARCH ARTICLE

Open Access

Melioidosis in Penh, Cambodia

Erika Vlieghe, Lim I
Chun Kham, Chhun H
Olivier Koole, Soph
and Jar

Pulmonary melioidosis in Cambodia: A prospective study

Blainde Rammaert¹, Julien Beauté¹, Laurence Borand¹, Sopheak Hem¹, Philippe Buchy¹, Sophie Goyet¹,
Rob Overtoom², Cécile Angebault¹, Vantha Te³, Patrich Lorn Try⁴, Charles Mayaud⁵, Sirenda Vong^{1*} and
Bertrand Guillard¹

We describe 58 adult patients (2007–2010). Diabetes was a risk factor for infections occurred during infection was present in 60%. 12% of all bloodstream infections were associated with melioidosis. Treatment.

Pleuropulmonary melioidosis in a Cambodian refugee

Charles K. Chan,* MD
Robert H. Hyland,* MD, FRCP[C]
Wolf D. Leers,† MD, PhD,
FRCP[C], Dip Bact
Michael A. Hutcheon,* MD,
FRCP[C]
Dorothy Chang,‡ MSc

A 47-year-old Cambodian refugee presented with an acute respiratory illness that featured consolidation of the lower lobe of the left lung and

Melioidosis, a common disease in Southeast Asia, is caused by *Pseudomonas pseudomallei*. We describe a patient from that endemic area who presented with acute pleuropulmonary melioidosis. The course of his illness illustrates the difficulties in microbiologic identification, the hazards of inadequate therapy, the ability of the organism to become resistant to many drugs, and the unusual involvement of the lower lobe of the left lung and adjacent pleura.

Positive soil samples for *B. pseudomallei*



40 samples collected from 10 rice fields along National Road 6 in Siem Reap during 2005-2006.

12 (30%) positive for *B. pseudomallei*

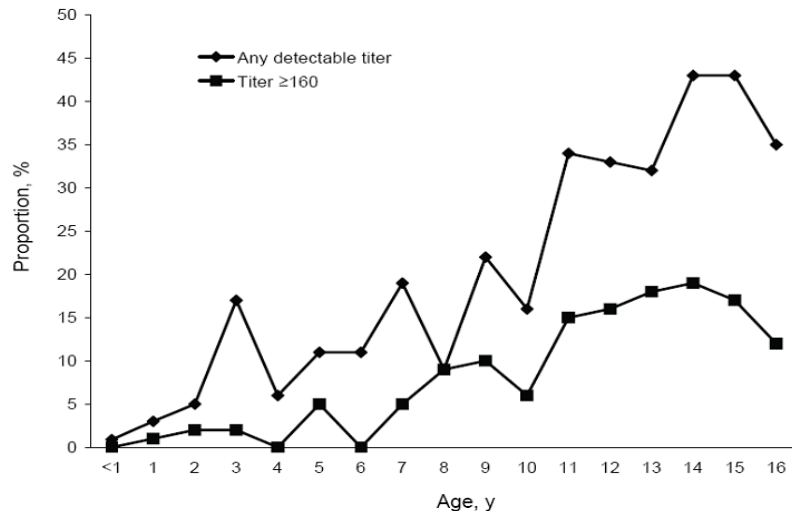
6 of 10 rice fields - at least 1 positive sample

- 1 rice field - positive for all 4 samples
- 3 fields - 2 positive samples
- 2 fields - 1 positive sample

Source: [Vanaporn Wuthiekanun](#),^{*} [Ngoun Pheaktra](#),[†] [Hor Putchhat](#),[†] [Lina Sin](#),[†] [Bun Sen](#),[†] [Varun Kumar](#),[†] [Sayan Langla](#),^{*} [Sharon J. Peacock](#),^{**} and [Nicholas P. Day](#)^{**} ***Burkholderia pseudomallei* Antibodies in Children, Cambodia** Emerg Infect Dis. 2008 February; 14(2): 301–303. doi: [10.3201/eid1402.070811](https://doi.org/10.3201/eid1402.070811)

SRP study 2006 – 16% children positive antibodies

The proportion of children with detectable *B.pseudomallei* AB titers increased with age



***Burkholderia pseudomallei* Antibodies in Children, Cambodia**

Vanaporn Wuthiekanun,* Ngoun Pheaktra,†
Hor Putchhat,† Lina Sin,† Bun Sen,†
Varun Kumar,† Sayan Langla,*
Sharon J. Peacock,*‡ and Nicholas P. Day*‡



Emerging Infectious Diseases • Vol. 14, No. 2, February 2008

Figure. Indirect hemagglutination assay (IHA) titer for 968 children living in Siem Reap, Cambodia. The number of children tested in each year group is shown. None of the children was known to have melioidosis. The number of children by age group follows: <1 y, 106 children; 1 y, 98 children; 2 y, 93 children; 3 y, 54 children; 4 y, 50 children; 5 y, 62 children; 6 y, 55 children; 7 y, 57 children; 8 y, 44 children; 9 y, 49 children; 10 y, 50 children; 11 y, 41 children; 12 y, 49 children; 13 y, 66 children; 14 y, 54 children; 15 y, 23 children; 16 y, 17 children.



Meeting Report

A Report from the Cambodia Training Event for Awareness of Melioidosis (C-TEAM), October 2017

Sotharith Bory ¹ , Frances Daily ^{2,*}, Gaetan Khim ², Joanne Letchford ², Srun Sok ³, Hero Kol ⁴,
Muy Seang Lak ⁴, Luciano Tuseo ⁵, Chan Vibol ⁵, Sopheap Oeng ² and Paul Turner ^{6,7} 

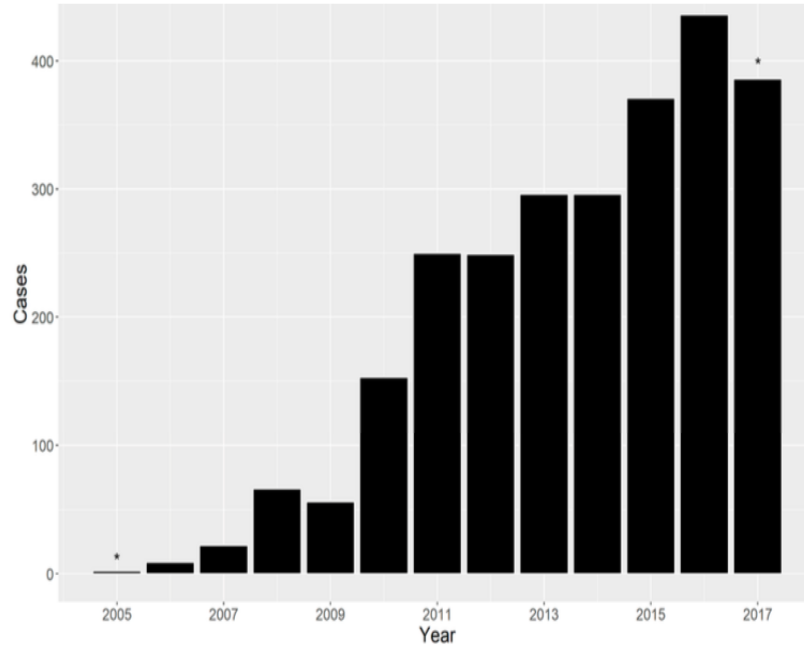


Figure 1. Annual numbers of confirmed melioidosis cases in Cambodia, October 2005–September 2017 (* incomplete years).

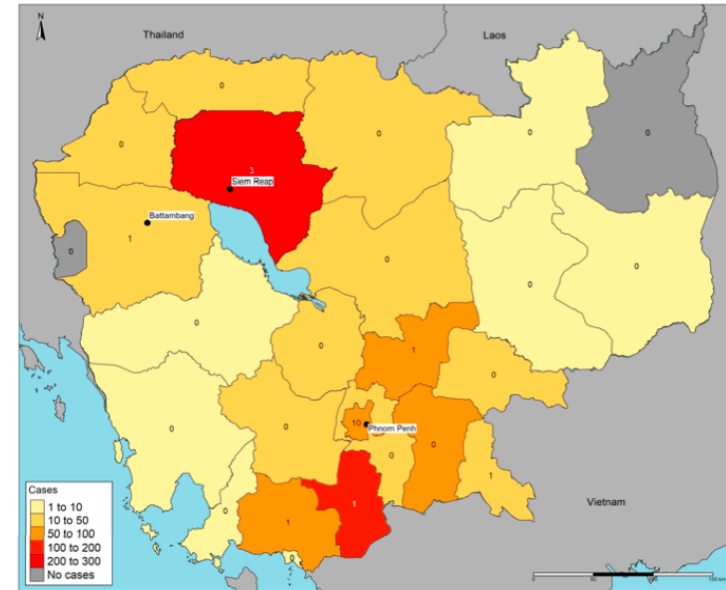
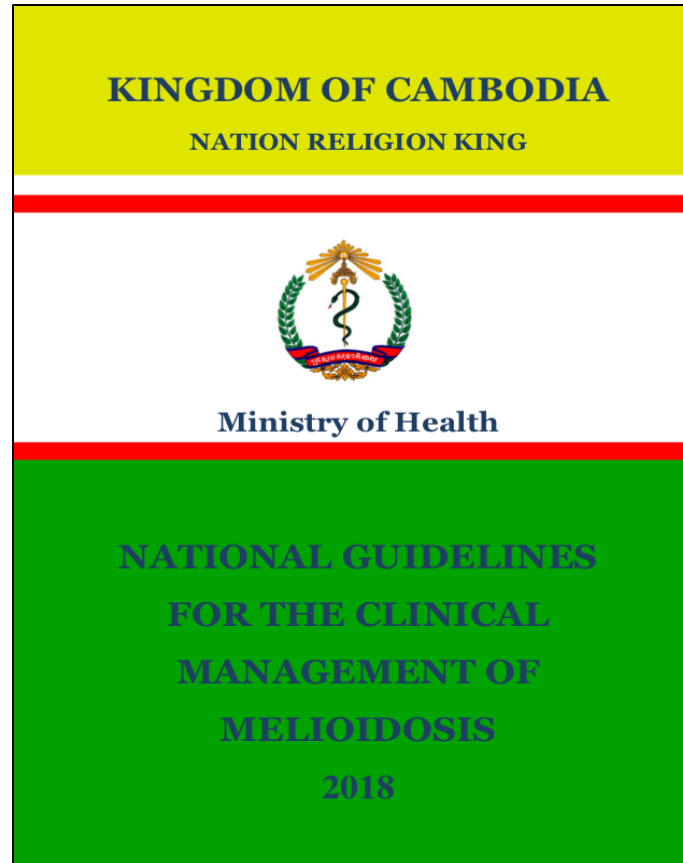


Figure 2. Geographic distribution of melioidosis cases in Cambodia, October 2005–September 2017. Province of residence was available in 889 confirmed cases. Shading represents the total number of culture-confirmed melioidosis cases per province. The numbers represent the count of participating microbiology laboratories per province; however, one was unable to contribute its culture confirmed cases prior to the meeting.

We have the answers here!



Conclusion

- ❖ Melioidosis has a high case fatality rate—up to 40% in some settings—especially when diagnosis and treatment are delayed. Chronic and relapsing forms also contribute to long-term disability and healthcare burden.
- ❖ Despite its impact, melioidosis has not been formally recognized as an NTD by the World Health Organization (WHO), leading to limited funding, research, and global surveillance efforts.

Mercie de votre Attention!