

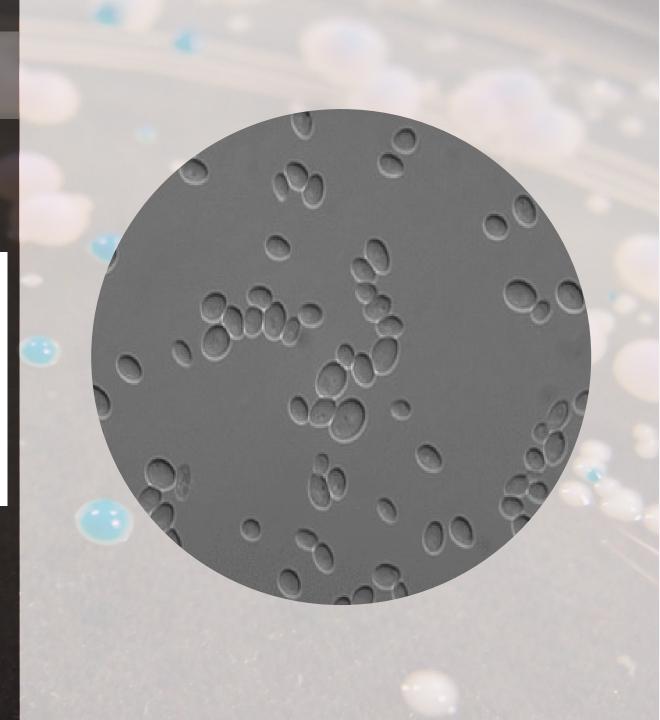
> Microbiol Immunol. 2009 Jan;53(1):41-4. doi: 10.1111/j.1348-0421.2008.00083.x.

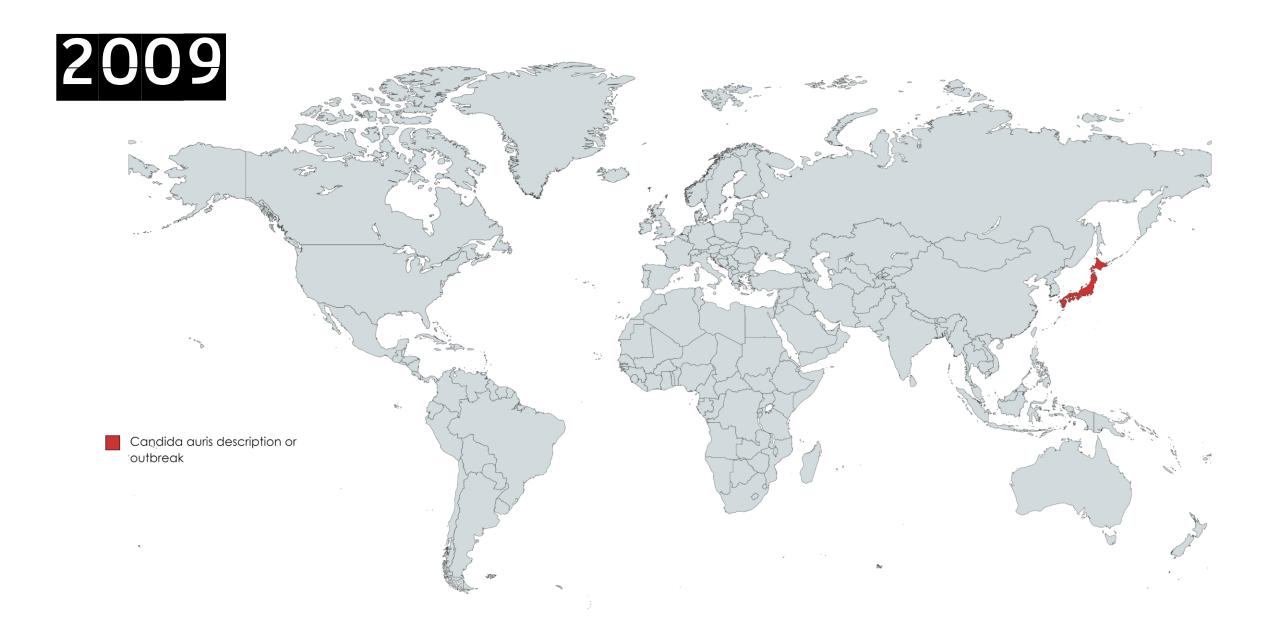
**ORIGINAL ARTICLE** 

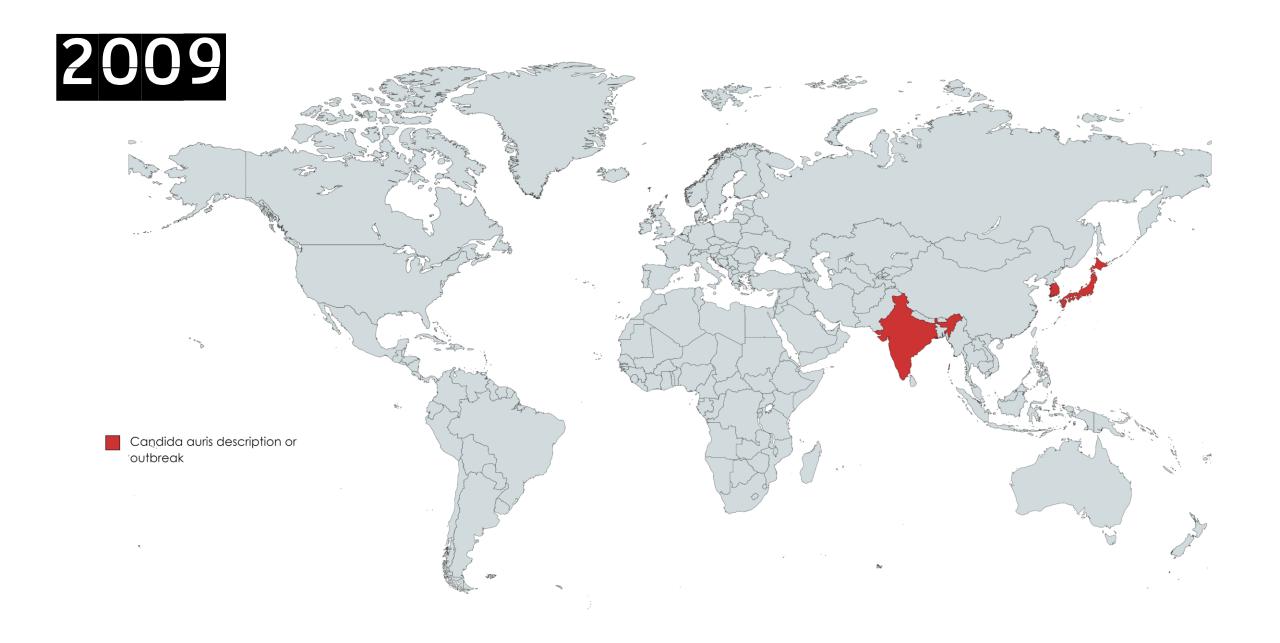
# Candida auris sp. nov., a novel ascomycetous yeast isolated from the external ear canal of an inpatient in a Japanese hospital

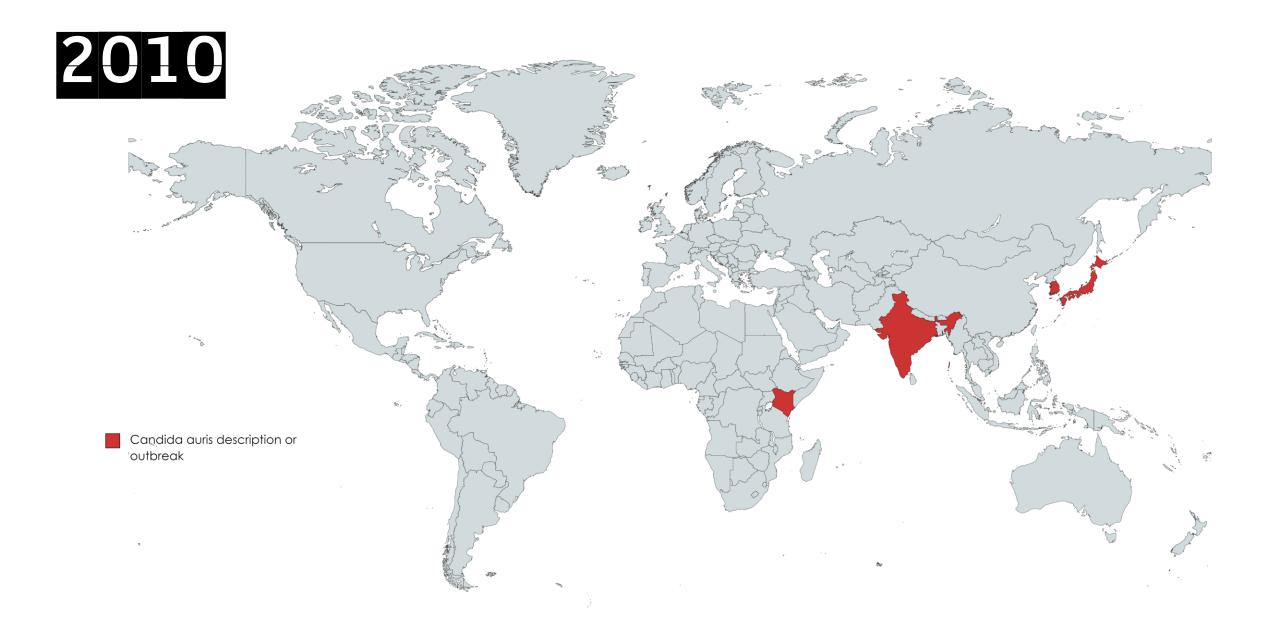
Kazuo Satoh<sup>1,2</sup>, Koichi Makimura<sup>1,3</sup>, Yayoi Hasumi<sup>1</sup>, Yayoi Nishiyama<sup>1</sup>, Katsuhisa Uchida<sup>1</sup> and Hideyo Yamaguchi<sup>1</sup>

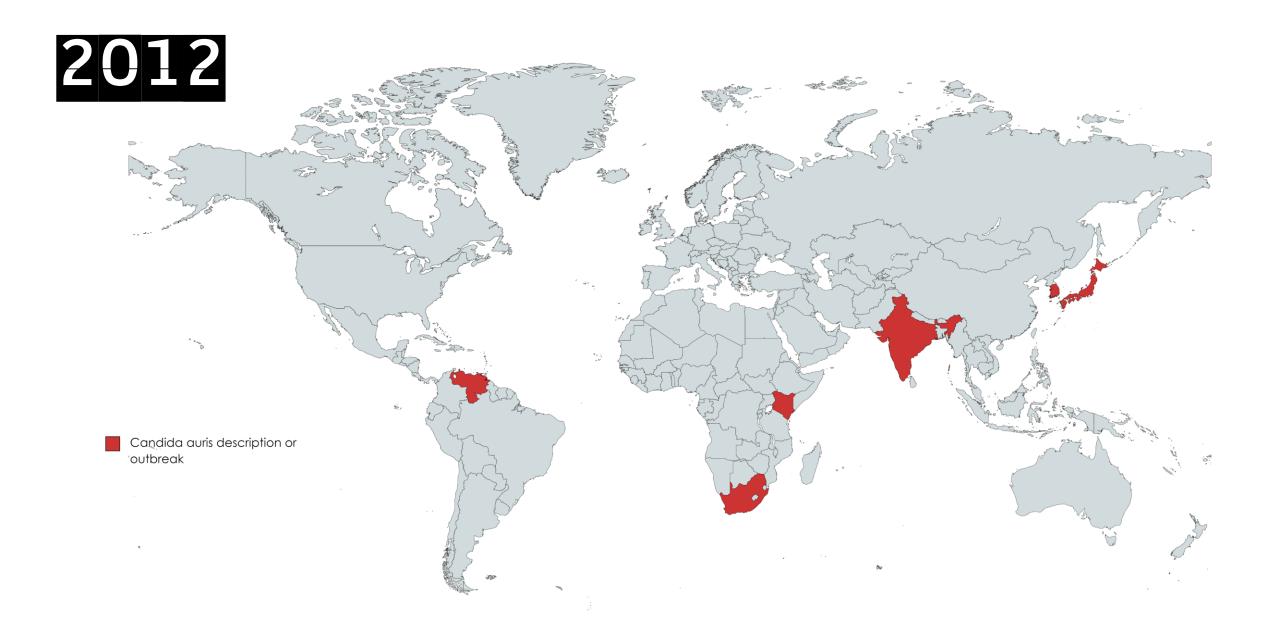
<sup>1</sup>Teikyo University Institute of Medical Mycology, 359 Otsuka, Hachioji, Tokyo 192-0395, <sup>2</sup>Japan Health Sciences Foundation, 13-4 Nihonbashi-Kodenmacho, Chuo-ku, Tokyo 103-0001 and <sup>3</sup>Genome Research Center, Graduate School of Medicine and Faculty of Medicine, Teikyo University, Otsuka 359, Hachioji, Tokyo 192-0395, Japan

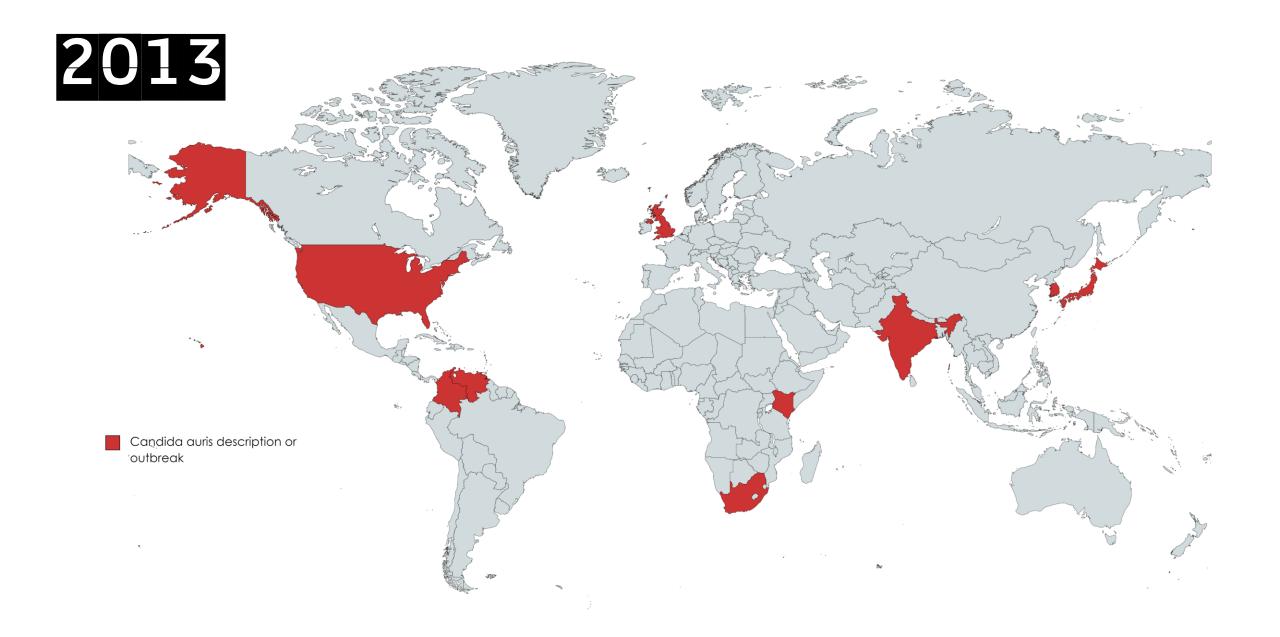


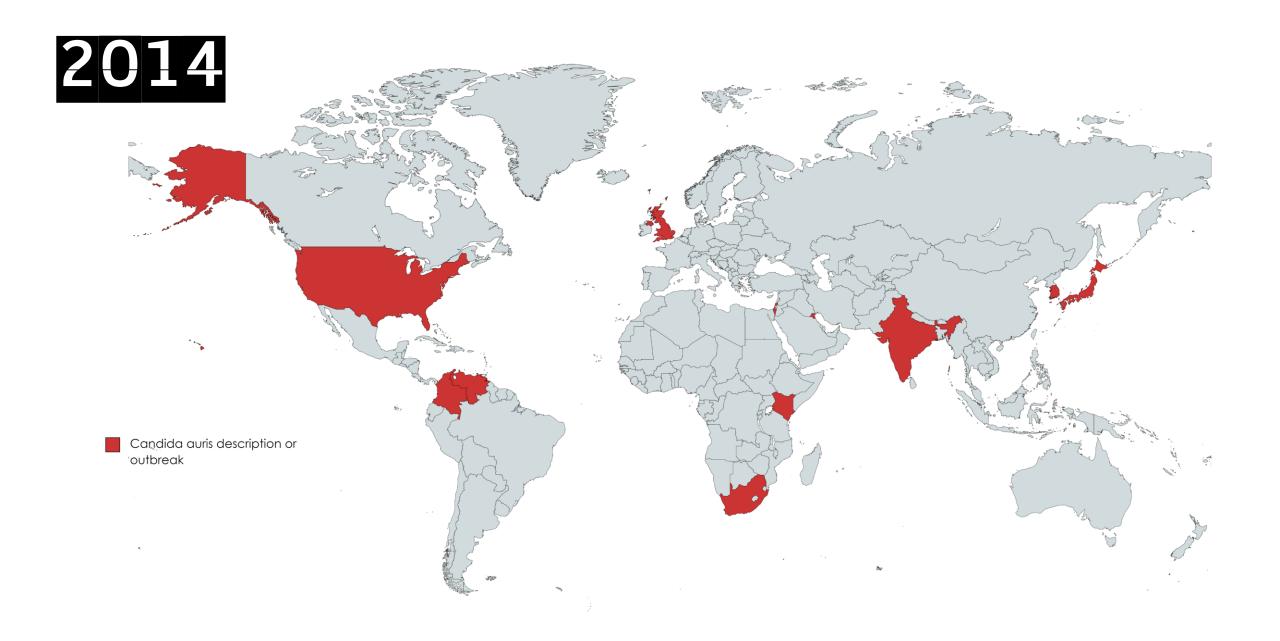


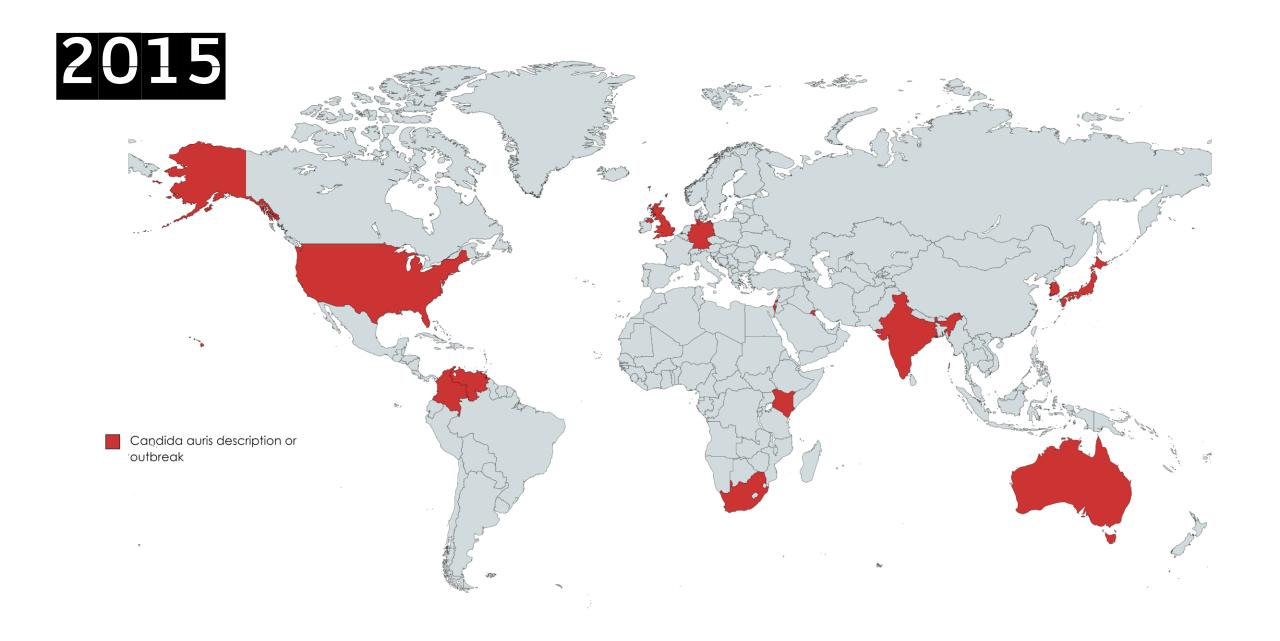


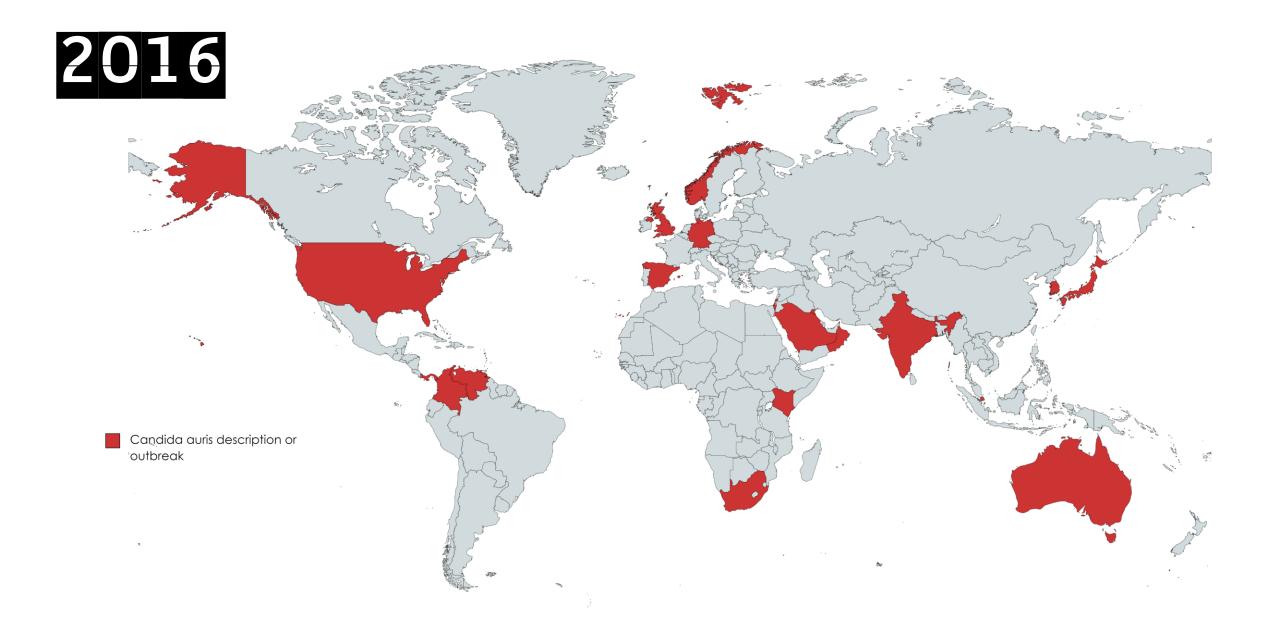
















#### CDC alerts in 2016:

Emergent multi-drug resistant fungal pathogen



## Clinical Alert to U.S. Healthcare Facilities - June 2016

#### Global Emergence of Invasive Infections Caused by the Multidrug-Resistant Yeast *Candida auris*

**Summary:** The Centers for Disease Control and Prevention (CDC) has received reports from international healthcare facilities that *Candida auris*, an emerging multidrug-resistant (MDR) yeast, is causing invasive healthcare-associated infections with high mortality. Some strains of *C. auris* have elevated minimum inhibitory concentrations (MICs) to the three major classes of antifungals, severely limiting treatment options. *C. auris* requires specialized methods for identification and could be misidentified as another yeast when relying on traditional biochemical methods. CDC is aware of one isolate of *C. auris* that was detected in the United States in 2013 as part of ongoing surveillance. Experience outside the United States suggests that *C. auris* has high potential to cause outbreaks in healthcare facilities. Given the occurrence of *C. auris* in nine countries on four continents since 2009, CDC is alerting U.S. healthcare facilities to be on the lookout for *C. auris* in patients.

### CDC alerts in 2016:

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- Severe nosocomial deep-seated infections



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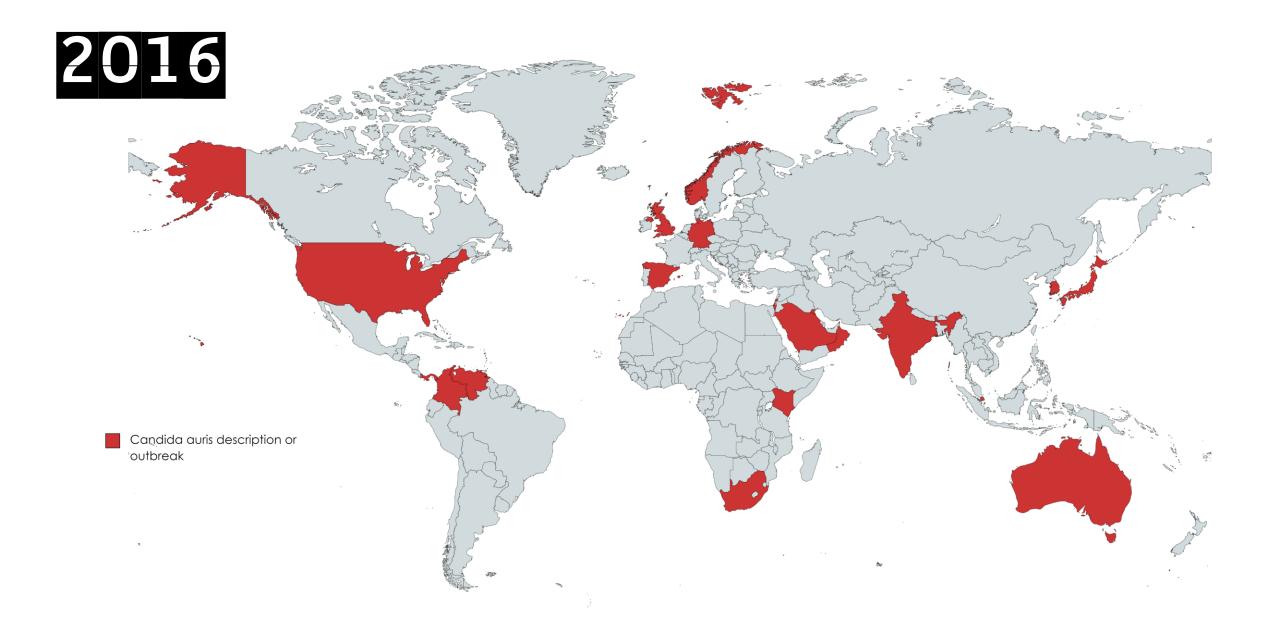
- Emergent multi-drug resistant fungal pathogen
- Severe nosocomial deep-seated infections
- High transmissibility and mortality → OUTBREAKS

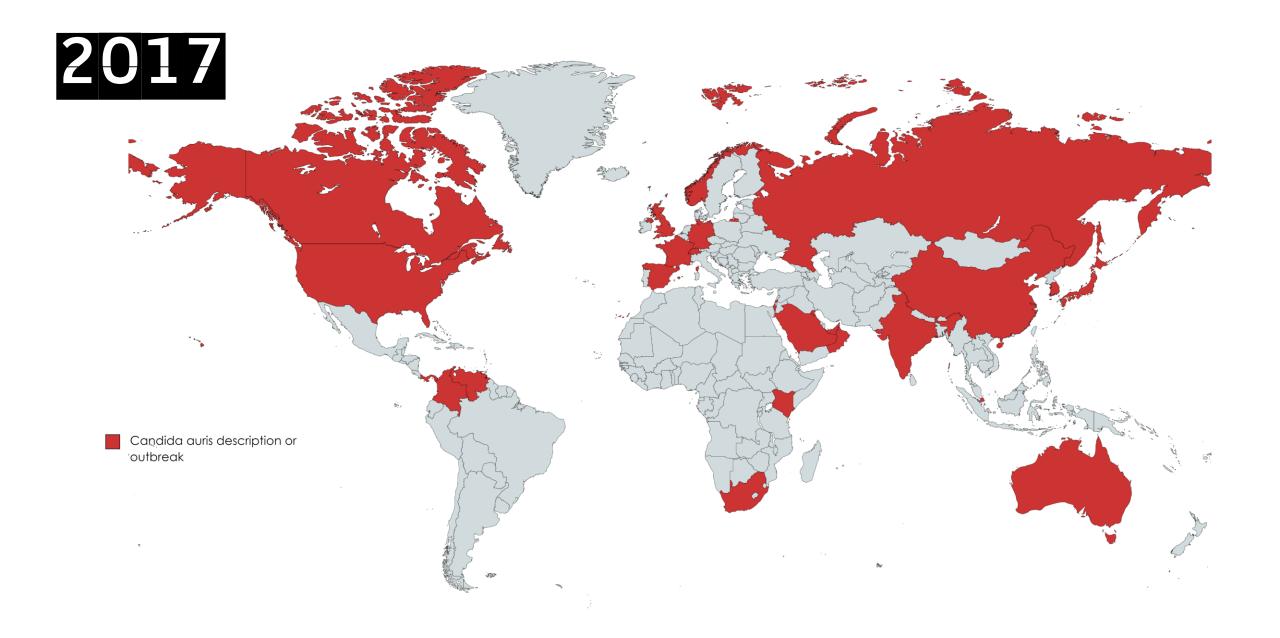


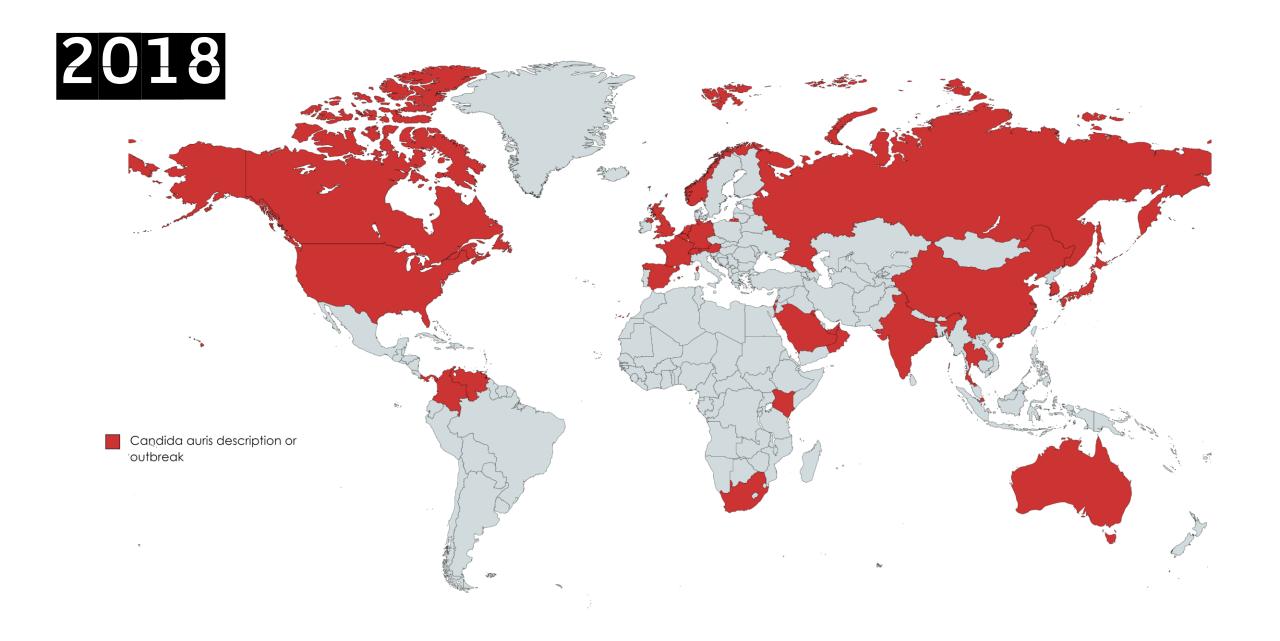
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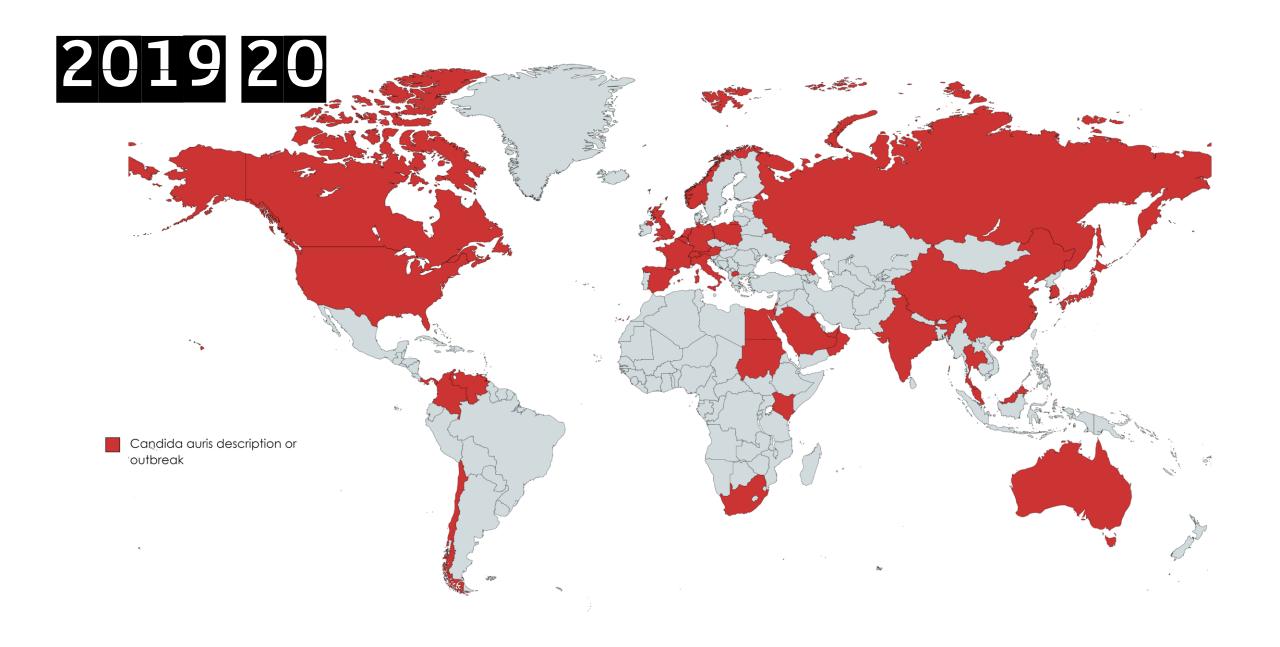
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### Global health threat:

- Emergent multi-drug resistant fungal pathogen
- Severe nosocomial deep-seated infections
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### ANTIBIOTIC RESISTANCE THREATS IN THE UNITED STATES



2019

#### **Urgent Threats**

- Carbapenem-resistant Acinetobacter
- Candida auris (C. auris)
- Clostridioides difficile (C. difficile)
- Carbapenem-resistant Enterobacteriaceae (CRE)
- Drug-resistant Neisseria gonorrhoeae (N. gonorrhoeae)

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- Increase with the COVID-19 pandemic

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## Candida auris: The new superbug on the block

By Lena Ciric **BBC News** 

(§ 17 August 2019)

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Jul 23, 2021, 01:51am EDT | 10.969 views

### Candida Auris Outbreaks: CDC Reports How This 'Superbug' Fungus Has Become More Dangerous



Bruce Y. Lee Senior Contributor ①

Healthcare

I am a writer, journalist, professor, systems modeler, computational and digital health expert, avocado-eater, and entrepreneur, not always in that order.





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**BBC News** 

Forbes

(1) 17 August 2019

### ANTIBIOTIC RESISTANCE THREATS IN THE UNITED STATES



**DEADLY GERMS, LOST CURES** 

# A Mysterious Infection, Spanning the Globe in a Climate of Secrecy

The rise of Candida auris embodies a serious and growing public health threat: drug-resistant germs.



**BBC News** 

(1) 17 August 2019

### ANTIBIOTIC RESISTANCE THREATS IN THE UNITED STATES



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### Outbreaks of Untreatable, Drug-Resistant Fungus Spread in 2 Cities

For the first time, the C.D.C. identified several cases of Candida auris that were resistant to all drugs, in two health facilities in Texas and a long-term care center in Washington, D.C.













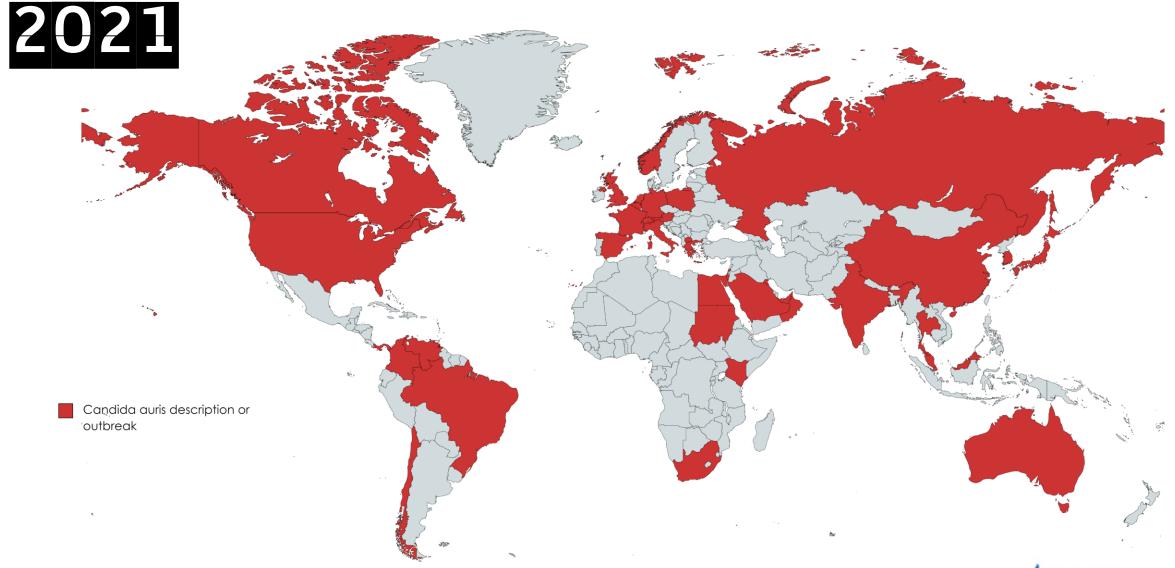




I am a writer, journalist, professor, systems modeler, computational and digital health expert, avocado-eater, and entrepreneur, not always in that order.

**BBC News** 

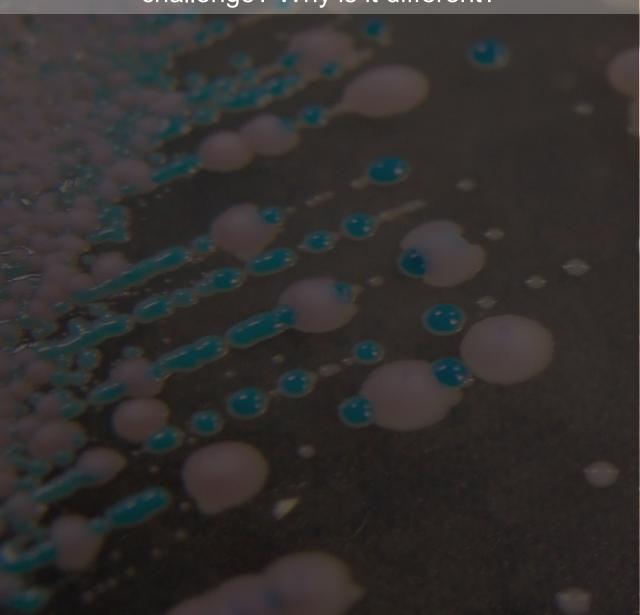
(1) 17 August 2019

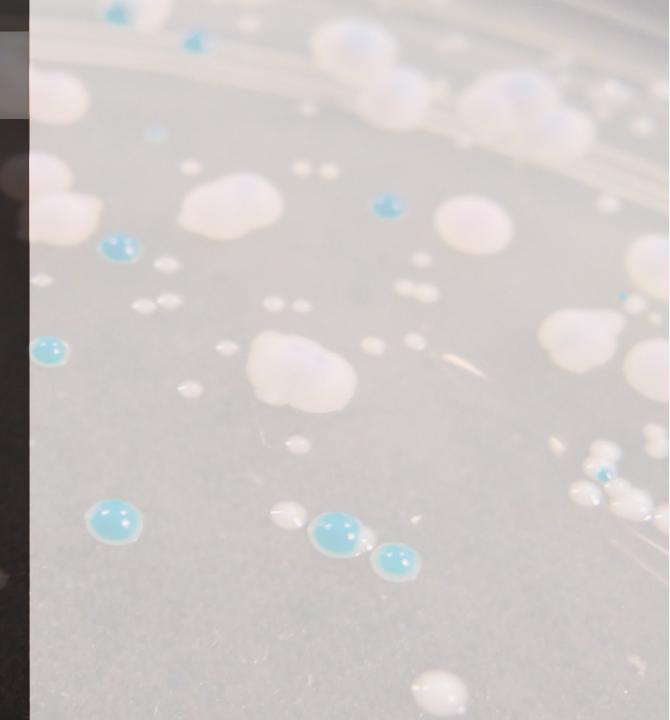


Countries from which *Candida auris* cases have been reported, as of February 15, 2021 This map is no longer being updated given how widespread *C. auris* has become.



Why does it represent a public health challenge? Why is it different?





# Why does it represent a public health challenge? Why is it different?

#### Morphogenetic plasticity

Different phenotypes – different behaviors

Dificulties in identification

#### Virulence

- Limited and diverse evidence
- Non-aggregative phenotypes more pathogenic
- Virulence factors

#### High mortality in susceptible patients

40-60% mortality of candidemia

High transmissibility and persistence

#### Extraordinary multi-drug resistance

Some isolates resistant to azoles, amphotericin and candines



#### MINIREVIEW

Therapeutics and Prevention July/August 2019 Volume 4 Issue 4 e00458-19 https://doi.org/10.1128/mSphere.00458-19

### Combined Antifungal Resistance and Biofilm Tolerance: the Global Threat of *Candida auris*

Ryan Kean<sup>a</sup>, Gordon Ramage<sup>b</sup>

Reference	Country (no. of isolates)	Method				400
(8)	India (350)	CLSI				100
(39)	United Kingdom <sup>a</sup> (119-128)	CLSI				
(72)	India* (123)	CLSI				80
(11)	India (90)	CLSI				
(14)	Colombia (87)	CLSI, Etest for AMB				%) s
(13)	United Kingdom# (73-79)	SYO				60 <del>g</del>
(32)	India (74)	CLSI				0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0
(10)	Spain (73)	EUCAST				40 is an
(2)	Korea# (61)	CLSI				40 sis
(9)	Kuwait (56)	Etest				· ·
(4)	Multiple <sup>b</sup> (54)	CLSI				20
(12)	Venezuela (18)	CLSI				
(1)	Korea (15)	CLSI, Etest for AMB				
			FLU	AMB	ECH	<b>0</b>

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			FLU	AMB	ECH	_0



- Globally four main clades
  - Possible clade V in Iran

Chow et al. Emerg Infect Dis. 2019 Sep;25(9):1780-1781.



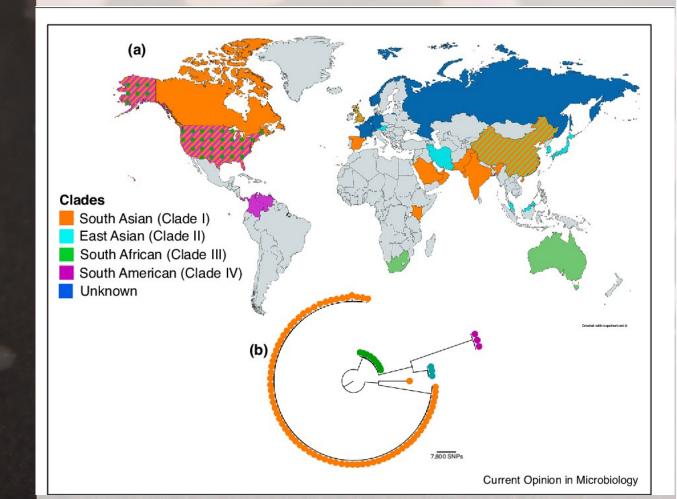
#### Current Opinion in Microbiology

Volume 52, December 2019, Pages 84-89



### Global epidemiology of emerging Candida auris

Johanna Rhodes, Matthew C Fisher M



- Globally four main clades
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No phylogenetical traceability

Simultaneous and independent appearance



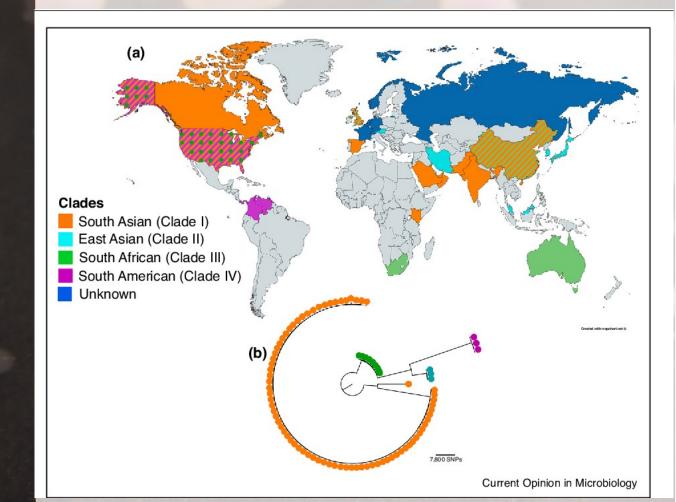
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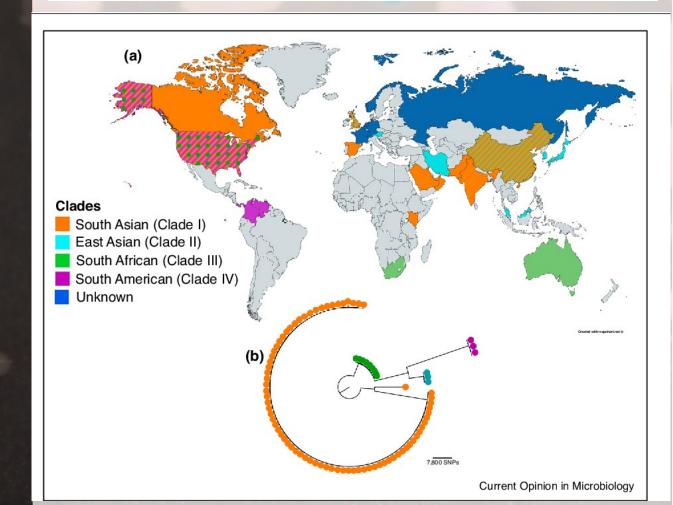




#### RESEARCH ARTICLE

Ecological and Evolutionary Science March/April 2020 Volume 11 Issue 2 e03364-19 https://doi.org/10.1128/mBio.03364-19

#### Tracing the Evolutionary History and Global Expansion of Candida auris Using Population Genomic Analyses



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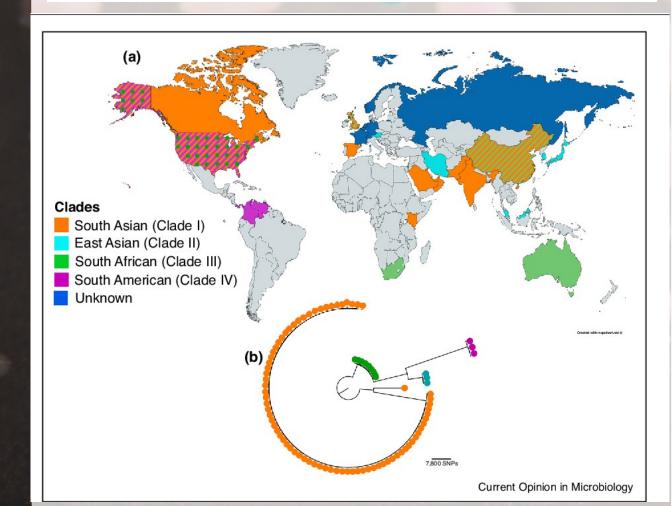




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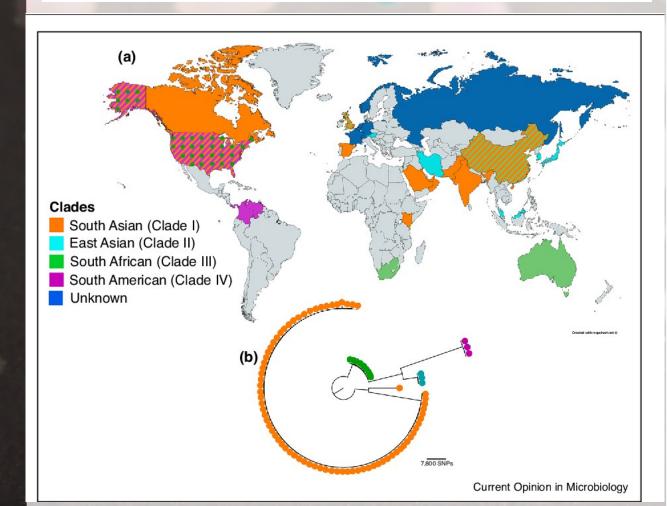




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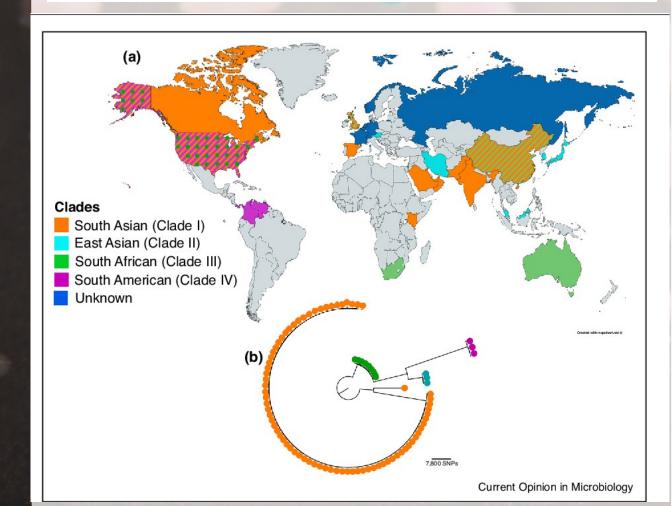




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#### What is the **Origin** of *C. auris?*

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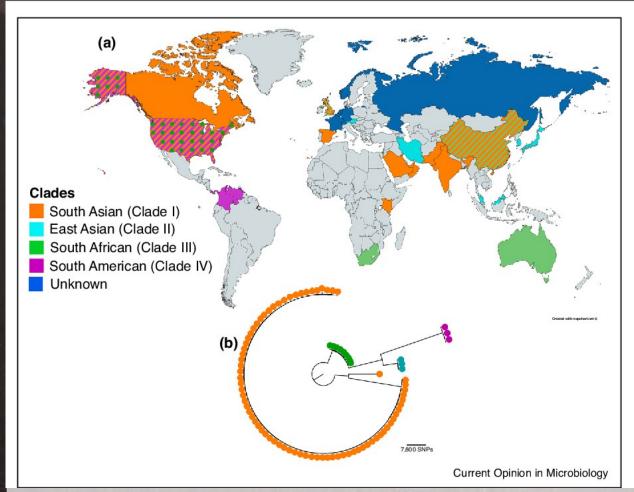




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- 148.000 fungal species described
- 5.1 million fungal species estimated



#### New scientific discoveries: Plants and fungi

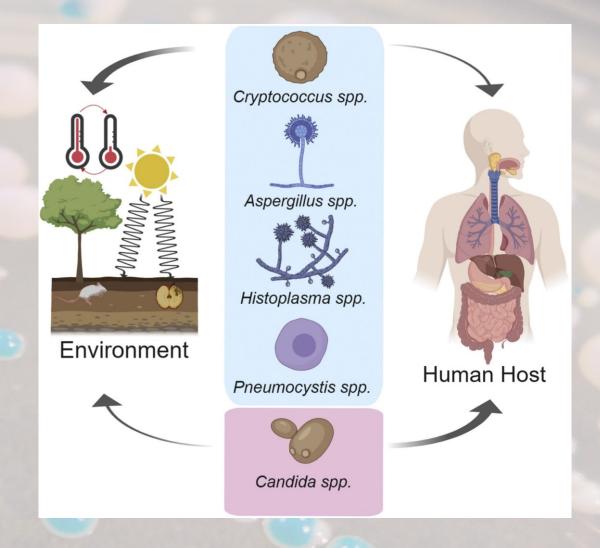
Martin Cheek ★, Eimear Nic Lughadha, Paul Kirk, Heather Lindon, Julia Carretero, Brian Looney ... See all authors ∨

> Am J Bot. 2011 Mar;98(3):426-38. doi: 10.3732/ajb.1000298. Epub 2011 Mar 2.

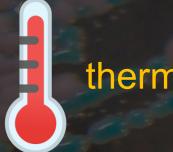
#### The fungi: 1, 2, 3 ... 5.1 million species?

Meredith Blackwell 1

- 148.000 fungal species described
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  - ↓ ↓ spp. infecting mammals and humans



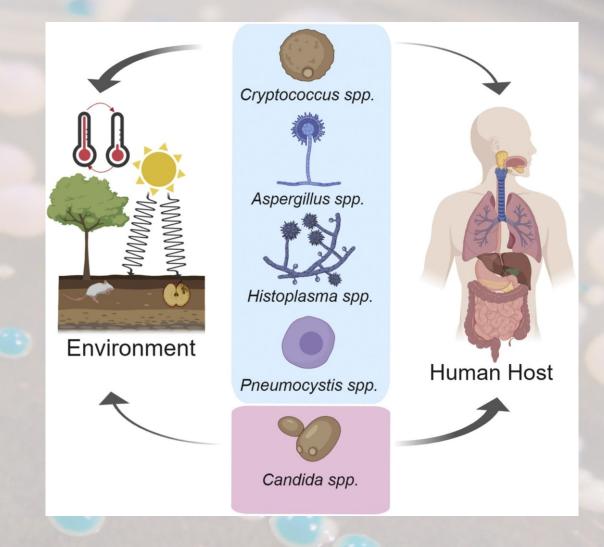
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thermal restriction barrier



adaptative immunity

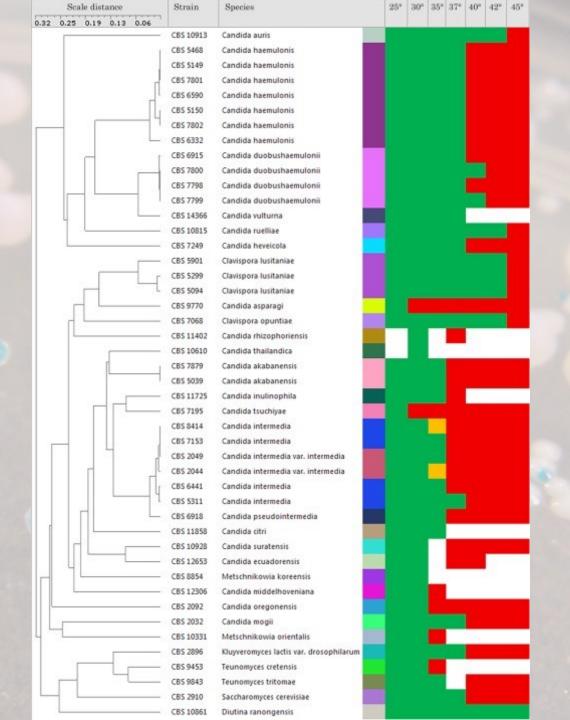


> PLoS Pathog. 2012;8(8):e1002808. doi: 10.1371/journal.ppat.1002808. Epub 2012 Aug 16.

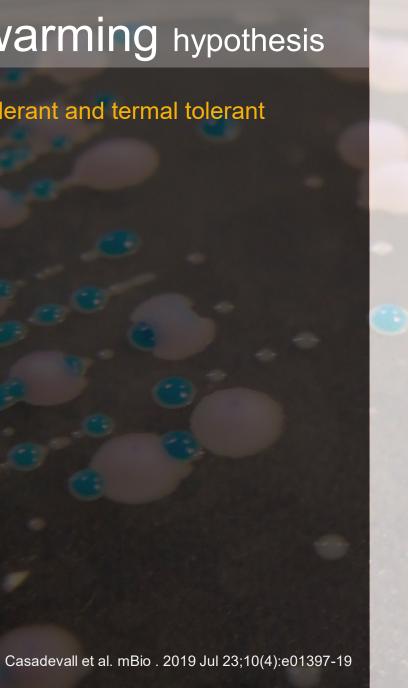
#### Fungi and the rise of mammals

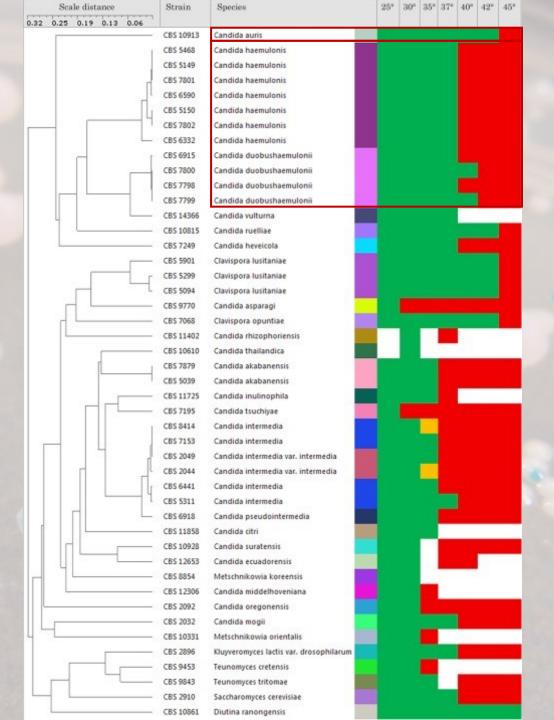
Arturo Casadevall 1

Casadevall et al. mBio . 2019 Jul 23;10(4):e01397-19



C. auris is halotolerant and termal tolerant

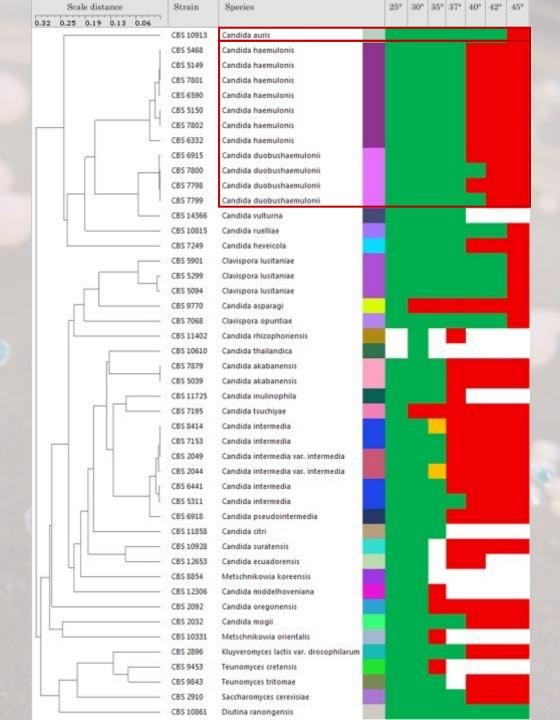




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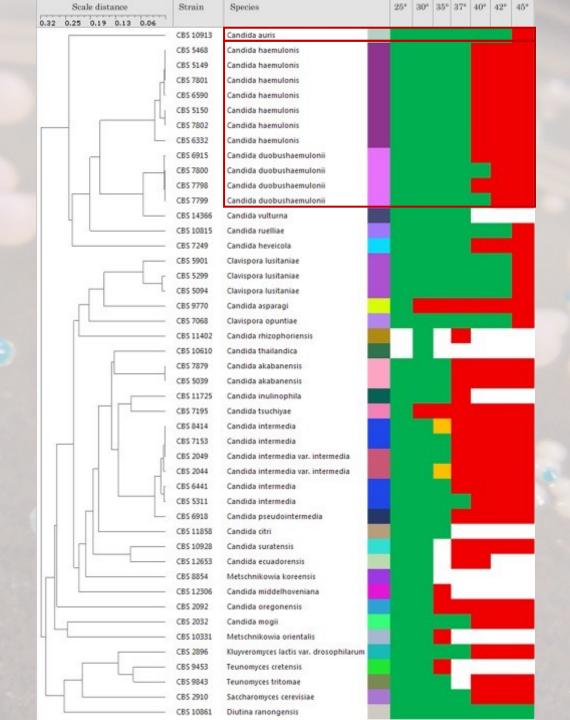
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Environmental Isolation of *Candida auris* from the Coastal Wetlands of Andaman Islands, India

Parth Arora<sup>a,b</sup>, Prerna Singh<sup>a</sup>, Yue Wang<sup>c</sup>, Anamika Yadav<sup>a</sup>, Kalpana Pawar<sup>a</sup>, Ashutosh Singh<sup>a</sup>, Gadi Padmavati<sup>b</sup>, Jianping Xu (b) c, and Anuradha Chowdhary (b) a

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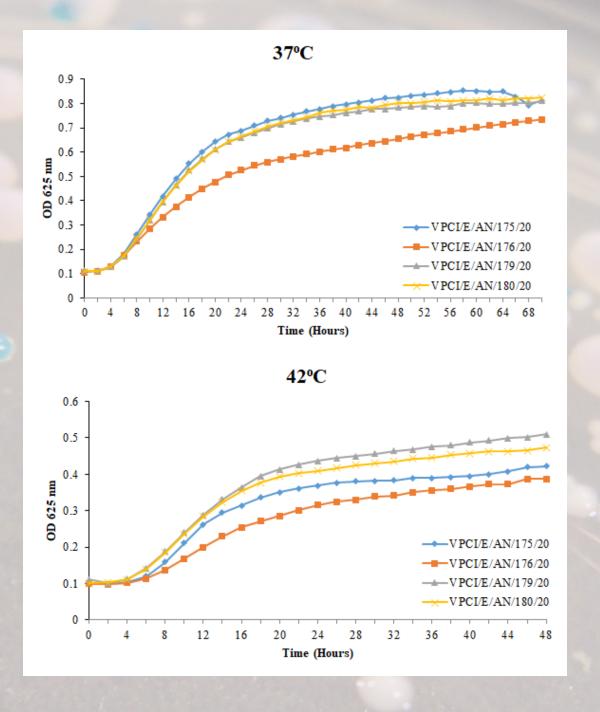


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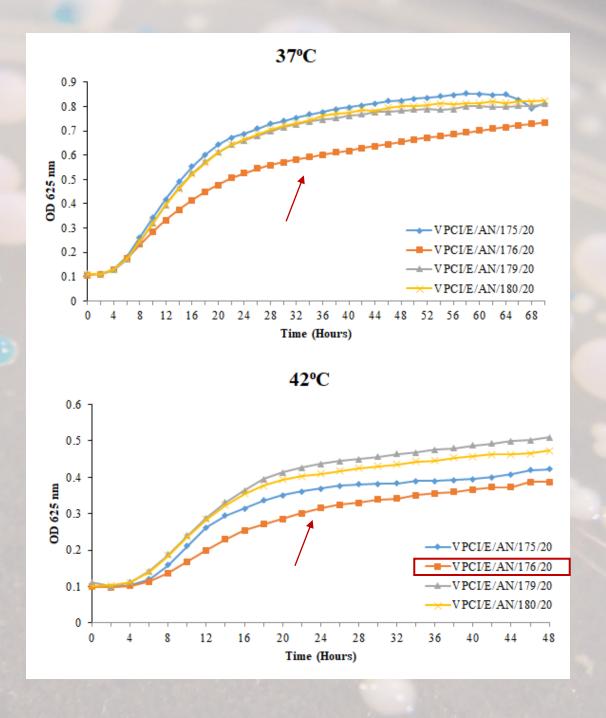


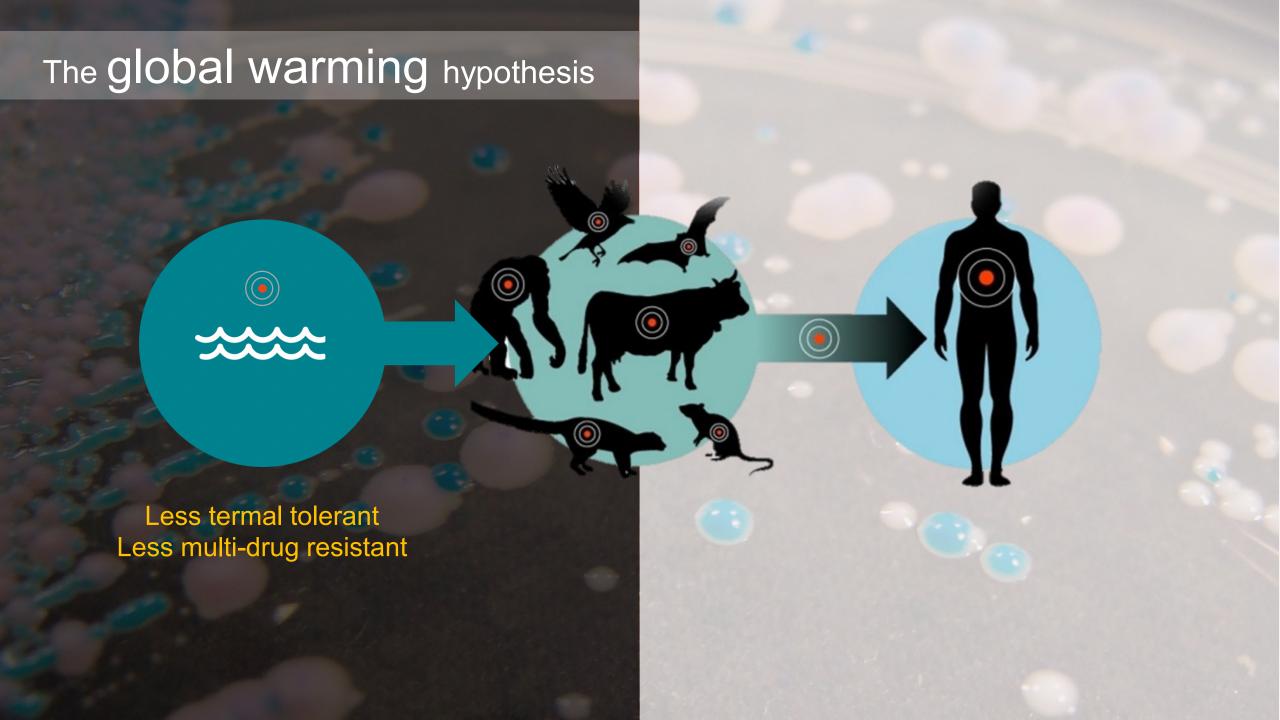
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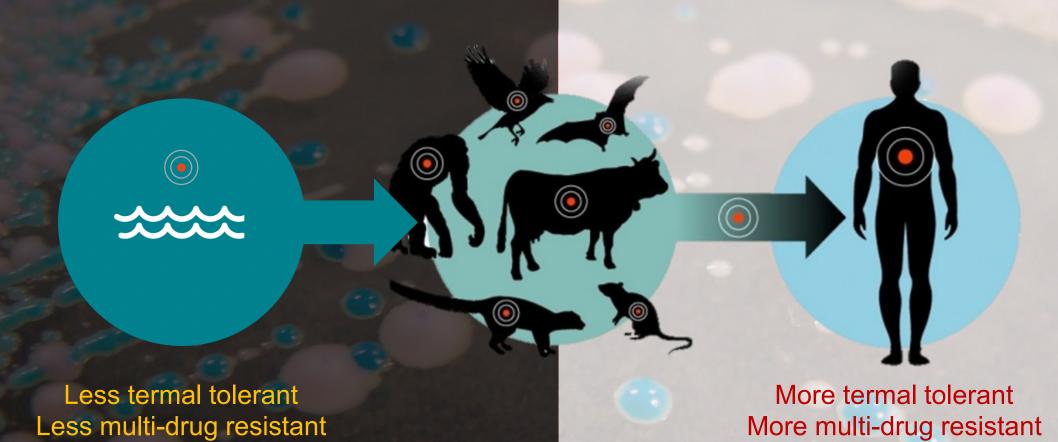


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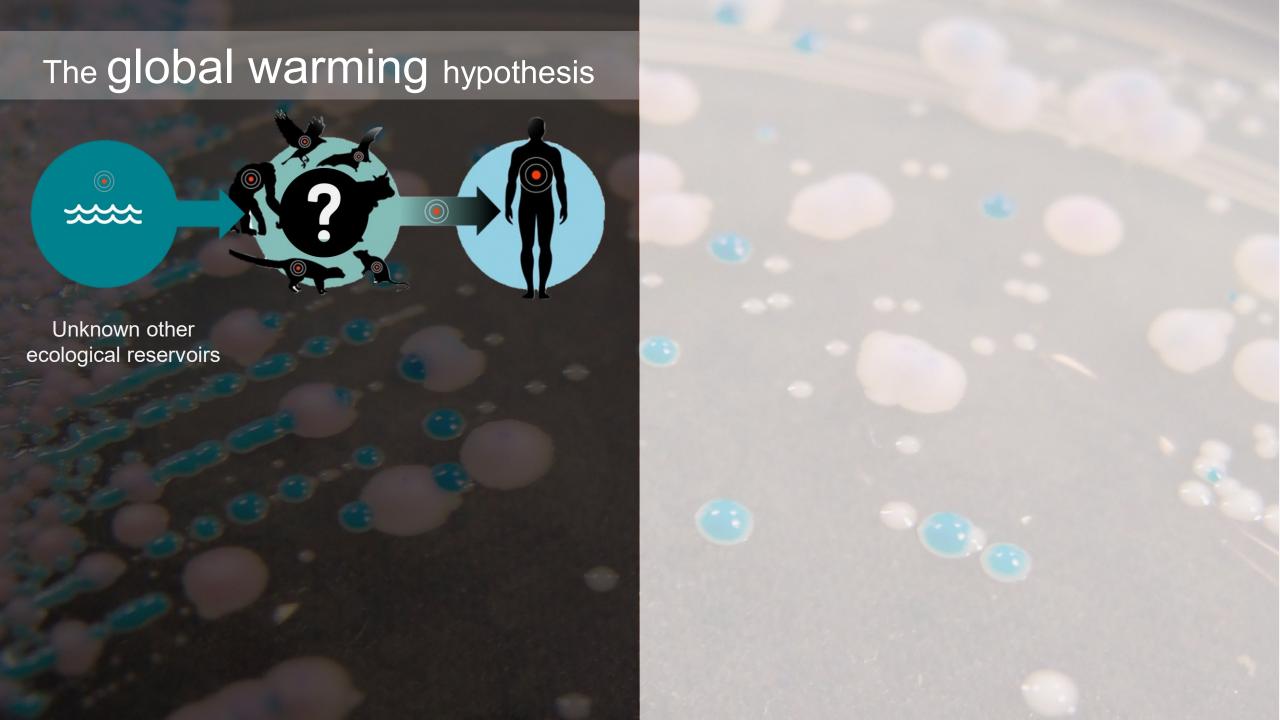


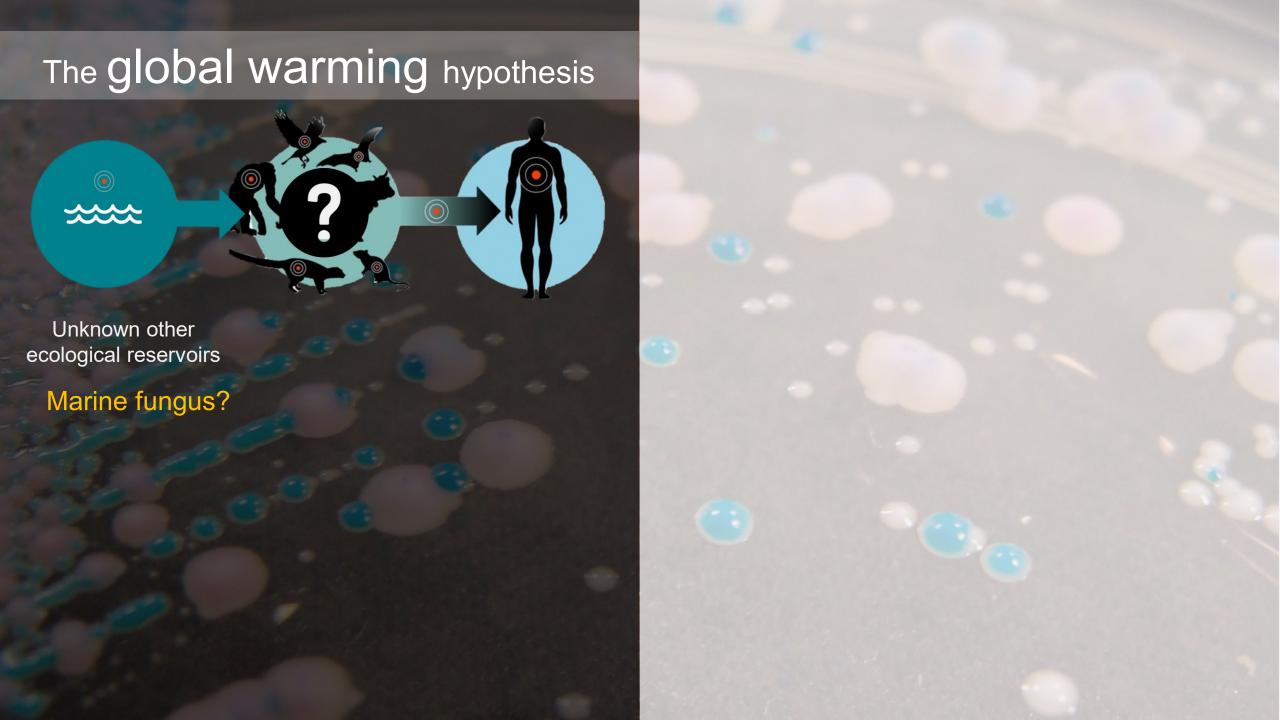






More multi-drug resistant







Unknown other ecological reservoirs

#### Marine fungus?

Osmotolerance

## On the Emergence of *Candida auris*: Climate Change, Azoles, Swamps, and Birds

Arturo Casadevall (b) a, Dimitrios P. Kontoyiannis (b) b, Vincent Robert<sup>c</sup>



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A clinical predictive model of candidaemia by *Candida auris* in previously colonized critically ill patients

V. Garcia-Bustos <sup>1,\*</sup>, M. Salavert <sup>2</sup>, A.C. Ruiz-Gaitán <sup>3</sup>, M.D. Cabañero-Navalon <sup>1</sup>, I.A. Sigona-Giangreco <sup>3</sup>, J. Pemán <sup>3</sup>



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Arturo Casadevall (b) a, Dimitrios P. Kontoyiannis (b) b, Vincent Robert<sup>c</sup>

#### Environmental Isolation of *Candida auris* from the Coastal Wetlands of Andaman Islands, India

Parth Arora<sup>a,b</sup>, Prerna Singh<sup>a</sup>, Yue Wang<sup>c</sup>, Anamika Yadav<sup>a</sup>, Kalpana Pawar<sup>a</sup>, Ashutosh Singh<sup>a</sup>, Gadi Padmavati<sup>b</sup>, Jianping Xu (b) c, and Anuradha Chowdhary (b) a

#### A clinical predictive model of candidaemia by *Candida auris* in previously colonized critically ill patients

V. Garcia-Bustos <sup>1,\*</sup>, M. Salavert <sup>2</sup>, A.C. Ruiz-Gaitán <sup>3</sup>, M.D. Cabañero-Navalon <sup>1</sup>, I.A. Sigona-Giangreco <sup>3</sup>, J. Pemán <sup>3</sup>

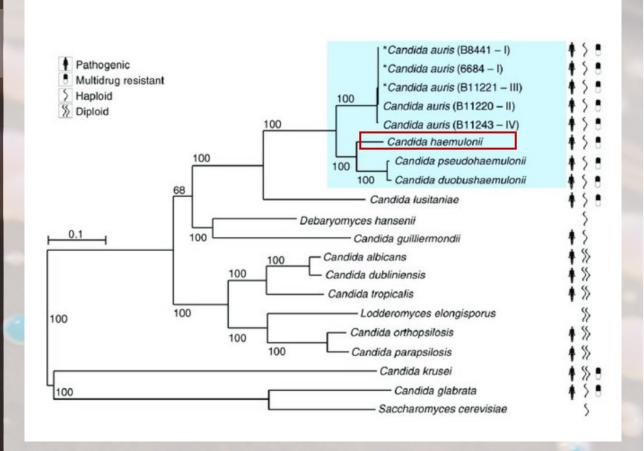




Unknown other ecological reservoirs

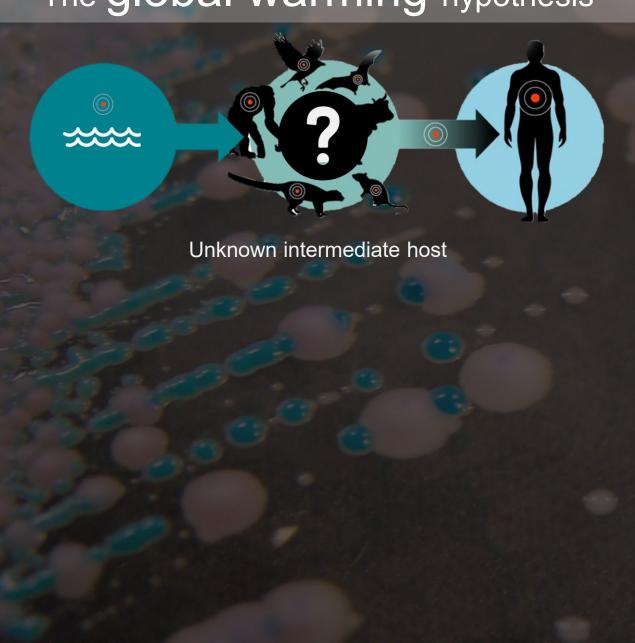
#### Marine fungus?

- Osmotolerance
- Andaman's findings
- Colonisation sources
- Coastal outbreaks
- Phylogeny clues

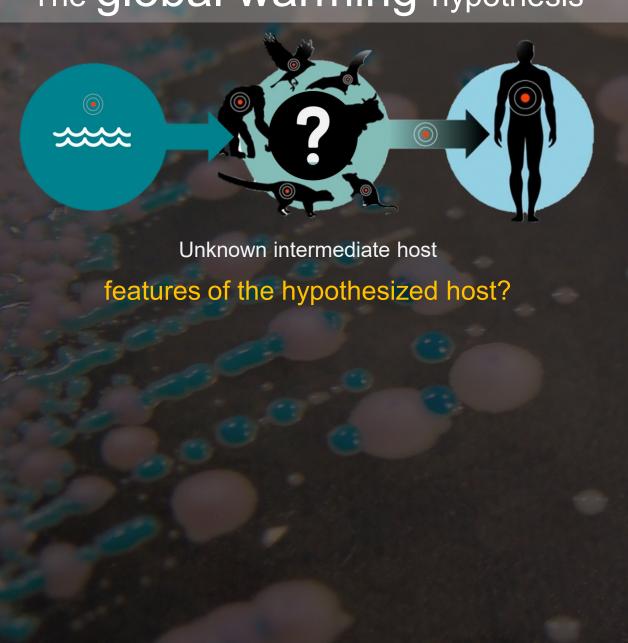


#### C. haemulonii isolated in:

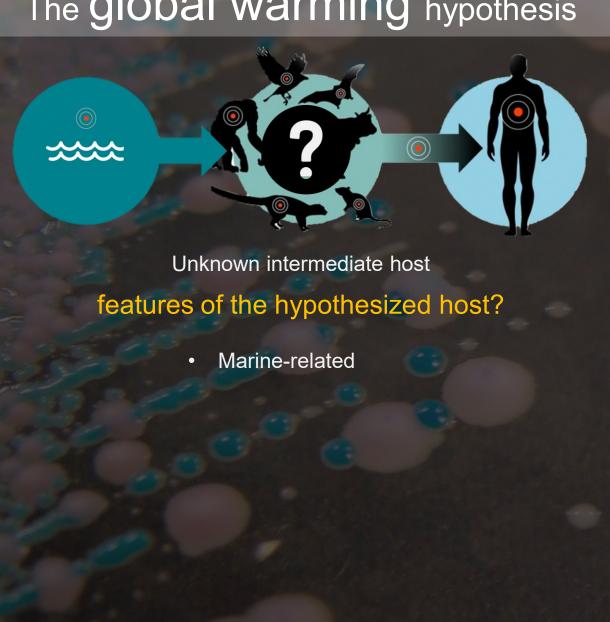
- Gut of Haemulon sciurus
- Portugal seawater (1962), Indian seawater (2011)
- Soft coral Palythoa variabilis, Brazil 2016
- Skin of a Dolphin (unknown species) in Suriname
- Pool with bottlenose dolphins (Tursiops truncatus)
- Aquaculture in shrimps















- Marine-related
- Migration routes



- Marine-related
- Migration routes
- Basal temperature ≤ human



- Marine-related
- Migration routes
- Basal temperature ≤ human
- Similar immunity



- Marine-related
- Migration routes
- Basal temperature ≤ human
- Similar immunity
- Documented colonisation



Unknown intermediate host

- Marine-related
- Migration routes
- Basal temperature ≤ human
- Similar immunity
- Documented colonisation
- Documented interspecies transmission



Unknown intermediate host

features of the hypothesized host?

- Marine-related
- Migration routes
- Basal temperature ≤ human
- Similar immunity
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- Documented interspecies transmission



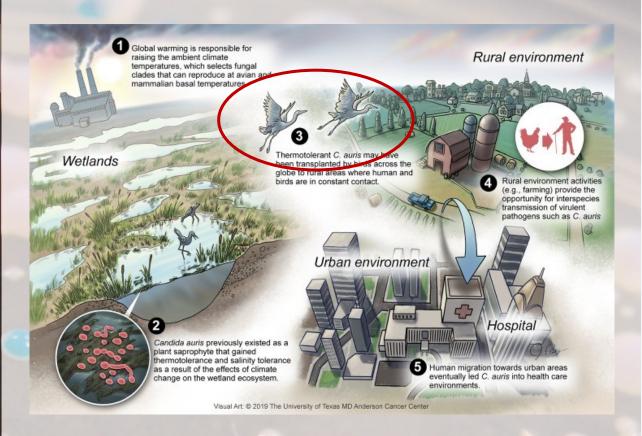
birds



Unknown intermediate host

features of the hypothesized host?

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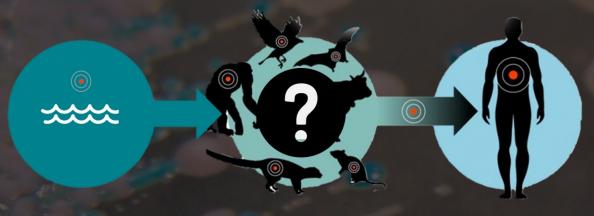


#### birds

Endocrinology of thermoregulation in birds in a changing climate

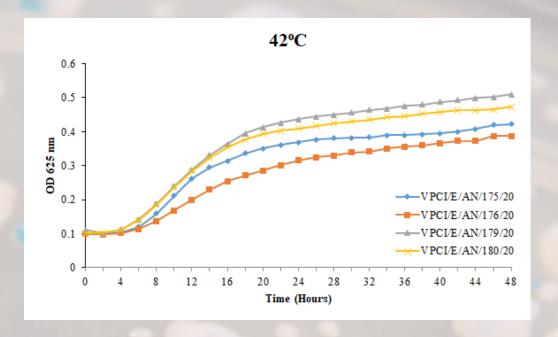
41.02 ± 1.29 °C

Suvi Ruuskanen <sup>a</sup>  $\stackrel{\triangle}{\sim}$  Bin-Yan Hsu <sup>a</sup>, Andreas Nord <sup>b</sup>



Unknown intermediate host

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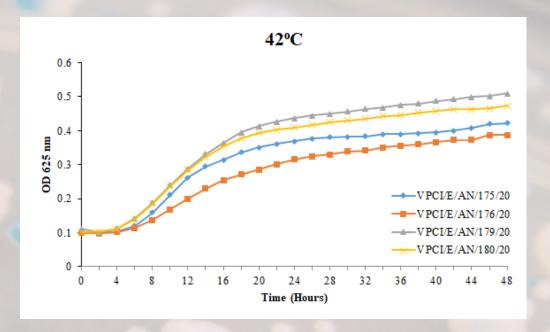




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- High evolutionary leap





#### RESEARCH ARTICLE

Ecological and Evolutionary Science March/April 2020 Volume 11 Issue 2 e03364-19 https://doi.org/10.1128/mBio.03364-19

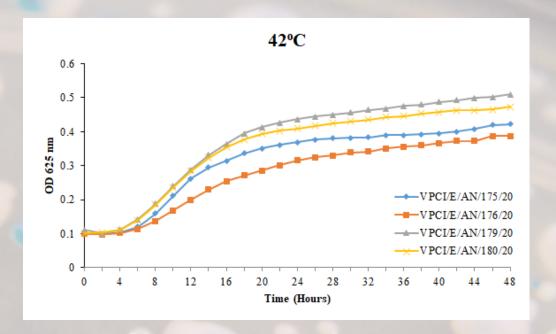
Tracing the Evolutionary History and Global Expansion of Candida auris Using Population Genomic Analyses



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From 380 – 36 years ago

Among clades: founder effect

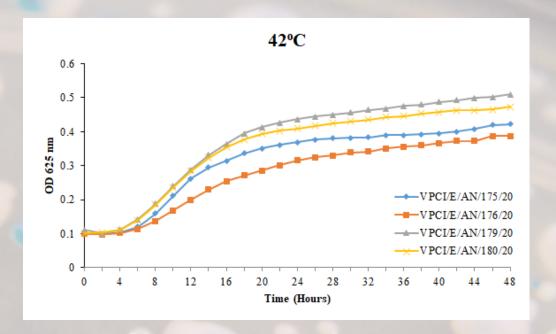




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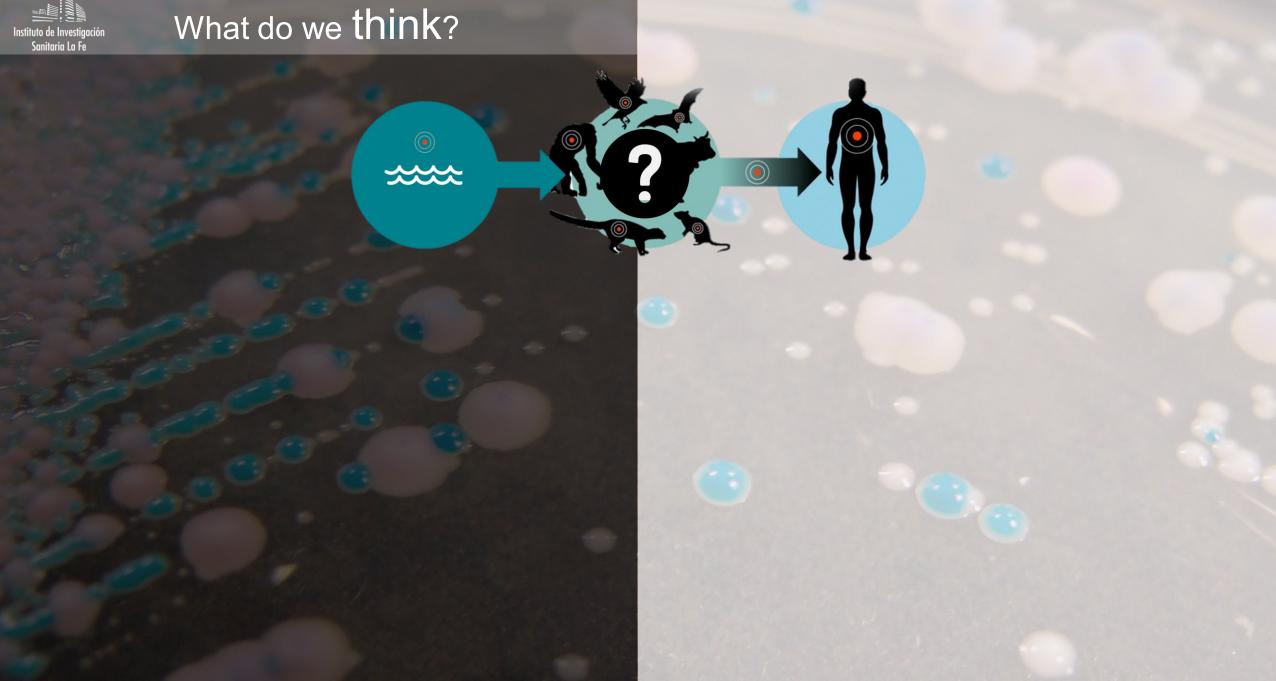
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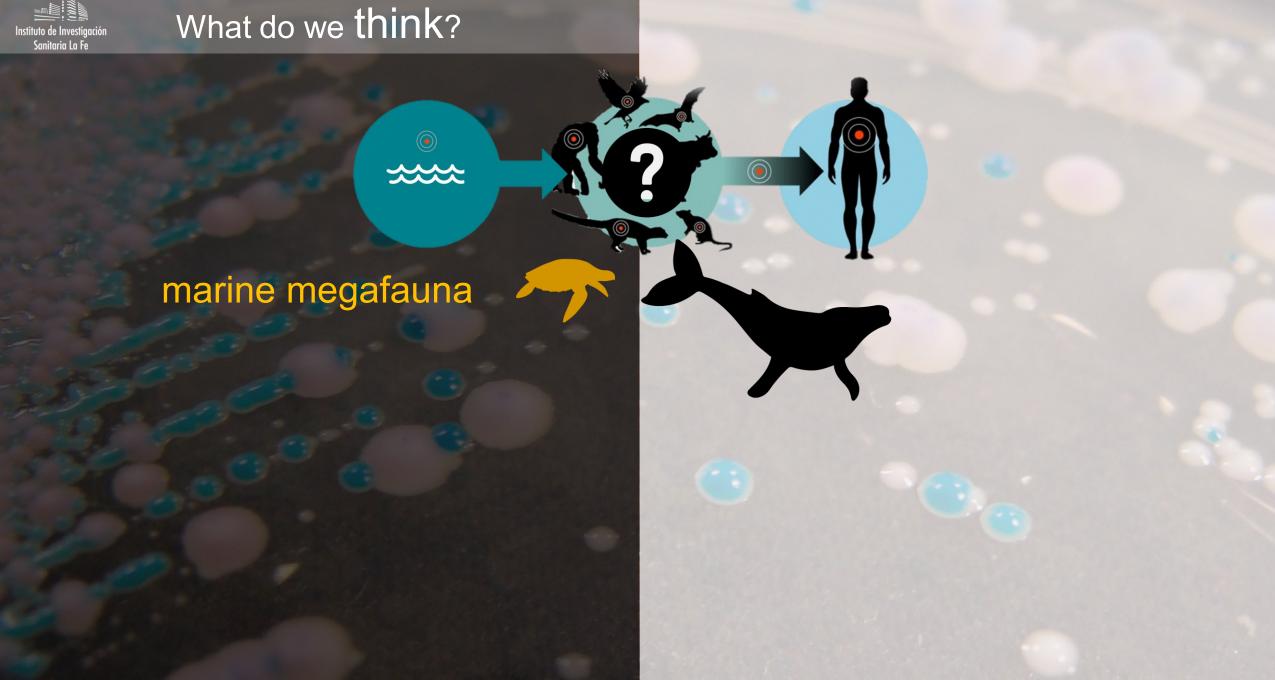
Among clades: founder effect













# What do we think?





- Marine-related
- Migration routes
- Basal temperature ≤ human
- Similar immunity
- Documented colonisation
- Documented interspecies transmission

















# What do we think?



- Marine-related
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Disseminated Candidiasis and Candidemia Caused by Candida palmioleophila in a Green Sea Turtle (Chelonia mydas)

Wen-Lin Wang 1,2, Pei-Lun Sun 3,400, Chi-Fei Kao 5,6, Wen-Ta Li 700, I-Jiunn Cheng 8 and Pin-Huan Yu 1,2,\*

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OCCURRENCE OF HUMAN-ASSOCIATED YEASTS IN THE FECES AND POOL WATERS OF CAPTIVE BOTTLENOSED DOLPHINS (Tursiops truncatus) 1 6

JOHN D. BUCK

#### Research Article

Candida albicans and C. tropicalis Isolates from the Expired Breathes of Captive Dolphins and Their **Environments in an Aquarium** 



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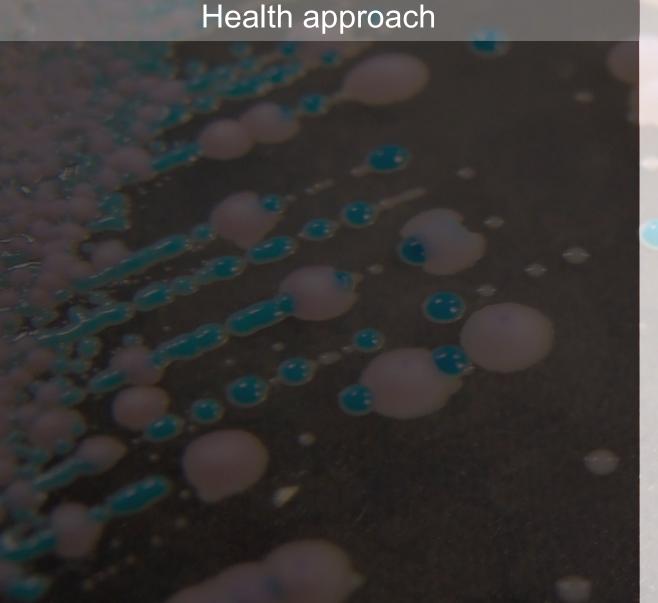
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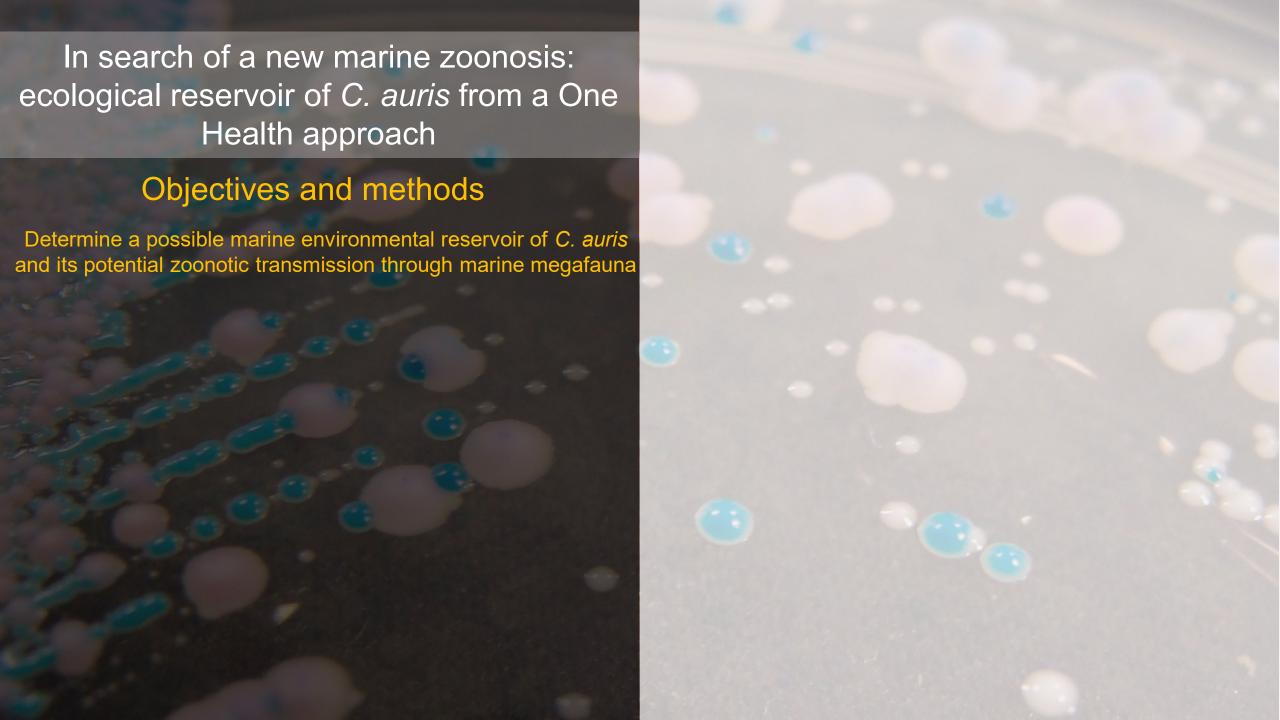
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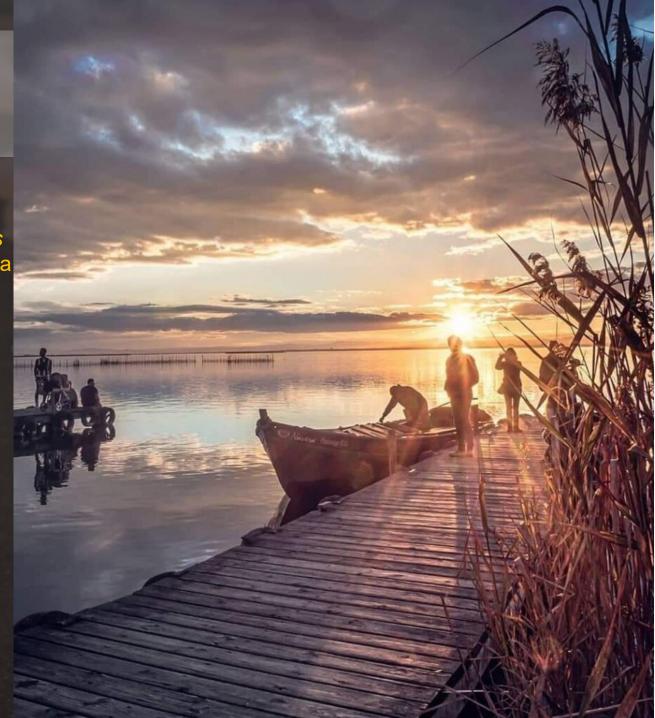




# Objectives and methods

Determine a possible marine environmental reservoir of *C. auris* and its potential zoonotic transmission through marine megafauna

• Detection of the presence of *C. auris* in salt marshes, continental freshwater, neritic and oceanic water and sediments, and supply and irrigation waters



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Departamento de Microbiología y Ecología



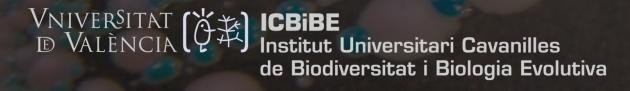
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# Objectives and methods

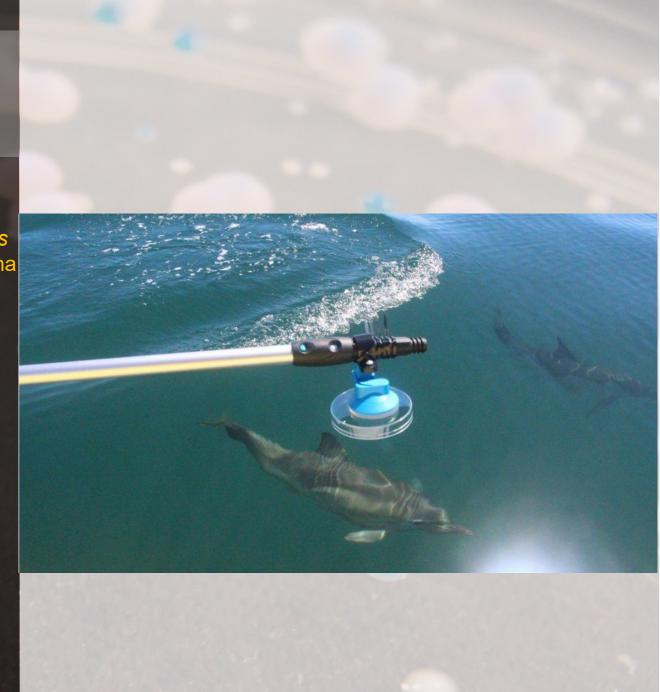
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## Objectives and methods

Microbiological identification

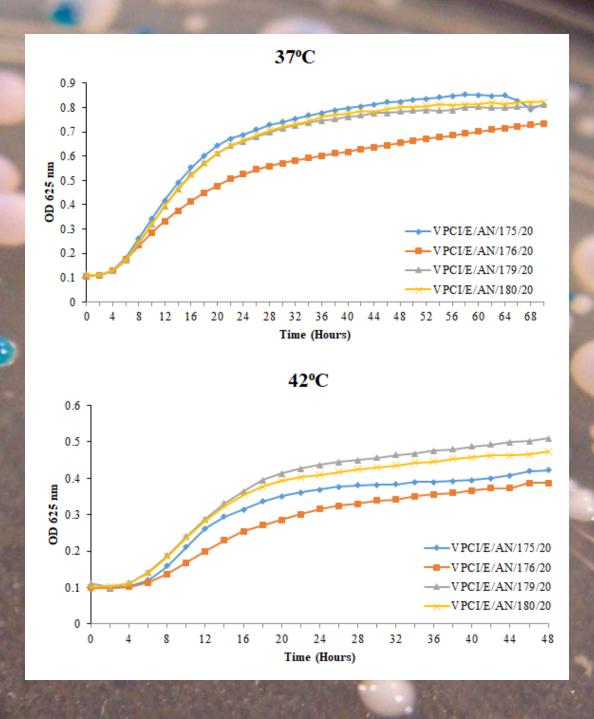
Culture (fungal and chromogenic media)



# Objectives and methods

Microbiological identification

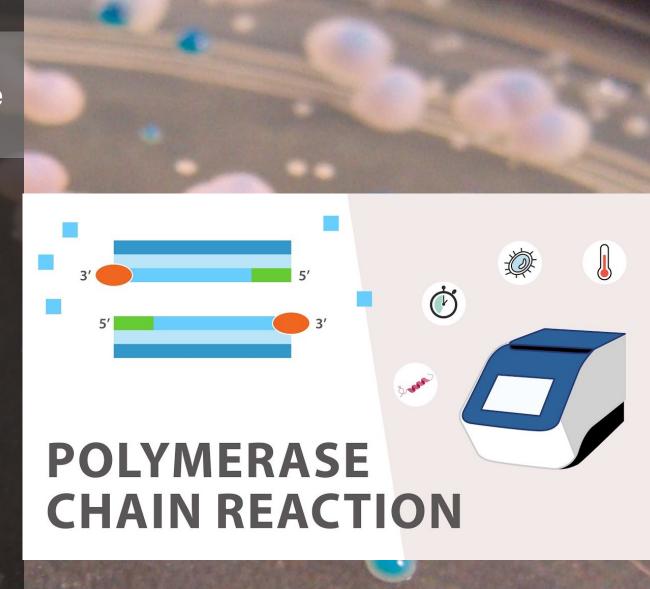
- Culture (fungal and chromogenic media)
  - Differential thermic growth assays



# Objectives and methods

Microbiological identification

- Culture (fungal and chromogenic media)
- PCR (screening)



# Objectives and methods

Microbiological identification

- Culture (fungal and chromogenic media)
- PCR (screening)
- Gene sequencing and phylogenetic analysis



