



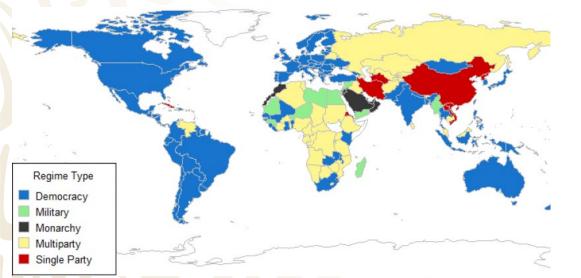
INSTITUTO DE DIAGNÓSTICO Y REFERENCIA EPIDEMIOLÓGICOS

Strengthening the International Cooperation against Emergencies between North America: Biosafety, Biosecurity and Epidemiological Surveillance of Infectious Diseases to the North Hemisphere

September 18th, 2019

countries in the world:

DE INDRE



 in Africa in Asia in Europe in America in Oceania



How big is the LATAM Region?

21 "Spanish speaking" countries in the world
7 in Central America
16 in the Caribbean region + 15 "dependent" territories
12 in South America + 3 "dependent" territories



What do we know about Mexico?

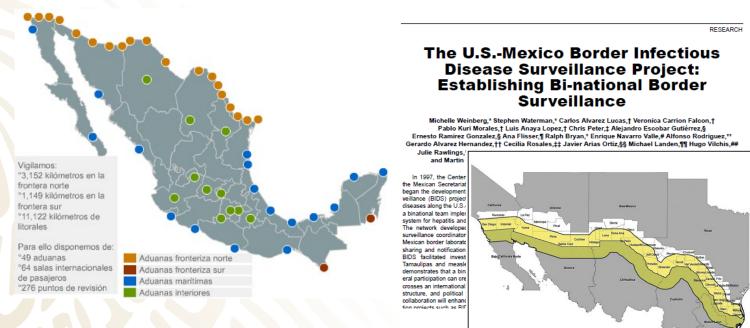


What do we know about Mexico?





How big is the challenge?





US-MEX International Border
US Border Awa
US Border Awa
US Border Reser Courties

States

0 37 575 150 225

Epidemiological Surveillance



What do we know about epidemiology efforts in the region?

PLOS ONE

RESEARCH ARTICLE

Zika virus: Epidemiological surveillance of the Mexican Institute of Social Security

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Abstract

Introduction

Methodology

confirmed cases

Conclusions

Introduction

Results

At the end of 2015, the first cases of Zika were identified in southern Mexico. During 2016, Zika spread as an outbreak to a large part of the country's coastal zones. The Zika epidemiological surveillance system records cases with clinical symptoms of Zika

virus disease (ZVD) and those confirmed by means of a reverse polymerase chain reaction

(BT-PCB) assay. This report includes the suspected and confirmed cases from 2016. Inci

dence rates were estimated by region and in pregnant women based on the proportion of

In total, 43,725 suspected cases of ZVD were reported. The overall incidence of suspected

were 4,168 pregnant women with suspected symptoms of ZVD, of which infection was con-

firmed in 1,082 (26%). The estimated incidence rate of ZVD for pregnant women nationwide

The incidence of Zika in Mexico is higher than that reported previously in the National Sys-

tem of Epidemiological Surveillance. Positive cases of Zika must be estimated and reported.

1/13

cases of ZVD was 82.0 per 100.000 individuals and 25.3 per 100.000 Zika cases. There

OPEN ACCESS

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Editor: Abdallah M, Samy, Faculty of Science, Ain Shams University (ASU), EGYPT

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Copyright: © 2019 Grajales-Muñiz et al. This is an open access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited

Data Availability Statement: All files are available from the public repository database. https:// figshare.com/articles/Zika virus epidemiologica surveillance of the Mexican Institute of Social

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Zika virus (ZIKV) was originally identified in a sentinel rhesus monkey in the Zika Forest of Uganda in 1947. The virus is a member of the family Flaviviridae, genus Flavivirus, and is mainly transmitted to humans by Aedes genus of mosquitoes [1]. The first recorded outbreak

was 186.1 positive Zika cases per 100.000 pregnant women.

PLOS ONE | https://doi.org/10.1371/journal.pone.0212114 February 11, 2019

Epidemiology and Infection

cambridge.org/hyg

Analysis of influenza data generated by four epidemiological surveillance laboratories in Mexico, 2010-2016

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Original Paper

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Key words: Infectious disease epidemiology; influenza: molecular biology

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Abstract

rates of morbidity and mortality. Thus, analysi logical surveillance systems has vital importance analysis was performed using data generated by the Mexican Social Security Institute between positivity, seasonality, treatment choices and va the vaccine according to its composition for eac enza subtypes and different age groups wer A/H1N1pdm09 (48.7%), influenza A/H3N2 (2 subtyped (11%) and influenza A/H1N1 (6.69 between November and March, and there were every 2 years. An inadequate use of oseltamivir cination status in general varied between 12.1 an provide current information about influenza in both operational case definitions and medical p

Investigación Biomédica del Noreste IMSS, Monterrey, N.I. **

Vicerrectoria de Ciencias de la Salud. Universidad de Moni

Pedro Garza Garcia, N.L., Mexico; "Unidad de Investigación

Centro Médico Nacional 'Ignacio García Téllez' IMSS, Méri

Vigilancia Epidemiológica (LAVE), Unidad Médica de Alta I

Yucatán, Mexico: ⁸Cátedra CONACvT, División de Investiga

Occidente IMSS, Guadalajara, Jal., Mexico: ⁹Laboratorio d

Molecular, Centro de Investigación Biomédica del Noreste Laboratorios de Vigilancia e Investigación Epidemiológica Evaluation and Reform of Mexican National **Epidemiological Surveillance System**

Roberto Tapia-Conver, MD, MPH, MSc, Pablo Kuri-Morales, MD, MSc, Luis González-Urbán, IE, and Elsa Sarti, MD, ScD

sidered which diseases should be

reported immediately or weekly

and three standardized formats

were defined: a Weekly Report

for a 2-month period. Problems

At a third national meeting.

the cutoff. Participants also conand whether a case study should sus through national meetings of epidemiologists, using a conceptual model be conducted. Listings were subof requirements, leadership, participation, and motivation. jected to a frequency analysis, The new SINAVE is run by committees that use data from 16 468 local health centers that generate homogeneous information from all health institutions. Indicators, flowcharts, and standardized instruments were creof New Cases of Disease (EPI-1ated. The reforms modernized SINAVE and strengthened epidemiologists' 95), a Case Study (EPI-2-95), leadership, consolidated local decision making, and assessed control acand an Outbreak Study (EPI-3tions needed to improve the health of the Mexican population. 95). A pilot test was conducted THE REFORM BEGAN WITH AN were discussed, and errors were

evaluation of SINAVE by means detected and corrected. of a qualitative interview, with a guideline to detect potential soluthe simple conceptual model of tions to several perceived prob-SINAVE reform, formats, manulems at the first national meeting als, and training program was of 150 state midamiologists responsed A Ginala Information. training plan A National Committee for Epidemiological Surveillance (CONAVE) was created through ministerial agreement to make surveillance statutory and comnulsory in the National Health System.4 CONAVE was conceived and designed according to the model of academic com mittees. With Mexico's complex health system, CONAVE has a unique value, because, for the first time, all organizations had been actively involved at SINAVE, and the Official Mexican Norm for Enidemiology Surveillance mandates that inunbournent (Element 1)?

an epidemiology bulletin, and a



The disease caused by the influenza virus is a use of antibiotics and antivirals.

To generate timely and reliable information for decision making in local health centers, Mexico's National Epidemiological Surveillance System (SINAVE) was evaluated and reformed. The reform was achieved by consen-

Representative pathogens identified in last 40 years (and counting....)

1972 Norovirus from diarrhea outbreak Norwalk Ohio 4 yrs. earlier 1994: Human herpesvirus 8=Kaposi's sarcoma-associated herpesvirus 1973 Rotavirus Major cause of infantile diarrhea worldwide 1994 Sabia virus Brazilian hemorrhagic fever 1975 Parvovirus B19 Fifth disease; Aplastic 1997; First Human outbreak of H5N1 1976 Cryptosporidium parvum Acute enterocolitis 1998: TT virus: a Transfusion Transmitted Hepatitis 1977 Ebola virus Ebola hemorrhagic fever 1998: Nipah virus: Encephalitis 1977 Legionella pneumophila Legionnaires' disease 1999: Anaplasma phagocytophilum (Human Granulocytic Ehrlichiosis/Anaplasmosis) 1977 Hantaan virus Hemorrhagic fever with renal syndrome (HFRS) 2000: Helicobacter canadensis another cause of diarrhea 1977 Campylobacter sp. Enteric pathogens distributed globally 2001: Human metapneumovirus (hMPV): Resp infections 1980 Human T-cell lymphoma leukemia lymphotropic virus-I (HTLV I) 2002: Corynebacterium appendicis: appendicitis 1981 Staphylococcus Toxic shock syndrome with toxin tampon use 2002: Dysgonomonas mossii :sepsis 1982 Escherichia coli, O157:H7 hemolytic uremic syndrome 2002: Kytococcus schroeteri: sepsis 1982 HTLV II Hairy cell leukemia 2003 SARS - CoV 1982 Borrelia burgdorferi Lyme disease 2003: "Bird flu" H5N1 new clad of birds & humans. 1983 Human immunodeficiency Syndrome-AIDS virus (HIV) 2004: Human cornoavirus NL63 (HCoV-NL63) 1983 Helicobacter pylori Gastric ulcer 2005: Porocephalus taiwana: a new pathogenic pentastomid 1988 Human herpesvirus-6 (HHV-6) Roseola subitum 2006: Prevotella baroniae septicemia and wound infection 1989 Ehrlichia chaffeensis Human ehrlichiosis 2007: New Orya-Fever like agent: Bartonella rochalimae sp. Nov 1989 Hepatitis C Parentally transmitted non-A, non-B hepatitis 2008 Lujo virus (Novel Arena virus of Africa) 1990 Recognition that Pneumocystis jiroveci was unique from P. carinii 2008 Mycobacterium chimaera (subtype of MAC) 1991 Guanarito virus Venezuelan hemorrhagic fever 2009 H1N1-2009 Pandemic 1991 Mycoplasma penetrans urogenital infection 2009 Wohlfahrtiimonas chitiniclastica sepsis 1992 Vibrio cholerae New strain associated with O139 epidemic cholera 2010 Negativicoccus succinicivorans 1992 Bartonella henselae Cat-scratch disease; bacillary angiomatosis 2011 Influenza A(H3N2)v virus 1992 Tropheryma whippelii Whipples Disease 2012 Middle East Respiratory Syndrome (MERS- CoV) 1992 Barmah Forest (BF) virus - Arbovirus Australia 2013 influenza A(H7N9) 1993 Hantavirus Hantavirus pulmonary syndrome isolates 2013 Severe fever with thrombocytopenia syndrome Virus (new bunyavirus) 1994: Asian Taeniasis: Human Tapeworm infection 2014 Avian influenza A H10N8



Epidemiological Surveillance: Mexico

- The Mexico Single Information System for Epidemiological Surveillance (SUIVE) created in 1981.
- SUIVE disseminates weekly bulletins detailing new cases of 142 diseases by subject, sex, and age group.
- The SUIVE Epidemiology Bulletin is the current bulletin for reporting morbidity data collected by National System of Epidemiological Surveillance (SINAVE).



Talking about epidemiological surveillance.....

PRO/AH/EDR> Rabies - Americas (44): USA, dog, skunk, raccoon, fox, human exp

RABIES - AMERICAS (44): USA, DOG, SKUNK, RACCOON, FOX, HUMAN EXPOSURE

A ProMED-mail post <<u>http://www.promedmail.org</u>> ProMED-mail is a program of the International Society for Infectious Diseases <<u>http://www.isid.org</u>>

In this post:

South Dakota/North Dakota: puppy, possible human exposure
 North Carolina: fox, human exposure

[3] New York: raccoon, human exposure

[4] Massachusetts: skunk, human exposure

[5] South Carolina: puppy, human exposure

[1] South Dakota/North Dakota: puppy, possible human exposure
 Date: Fri 30 Aug 2019
 Source: Jamestown Sun [edited]
 https://www.jamestownsun.com/lifestyle/pets/4639568-After-puppy-tests-positive-for-rabies-health-departments-in-the-Dakotas-seek-owners-of-littermates-



Situational diagnosis of biosafety and biosecurity in LATAM



Situational diagnosis: biosafety and biosecurity

Lack of:

- a) Financial resources
- b) Training
- c) Personnel assigned to the "biosafety and/or biosecurity" activities
- d) Interest in the topic
- e) Opportunities to "learn more about the topic"



What is affecting us?

- Unknown outbreaks
- Terrorism
- Activities without risk assessment
- Corruption
- Unemployment
- Migration challenges
- Excessive use of GMOs
- Use of unknown technology
- Others.....











What are governments doing right now?

How is communication transmitted to the scientists, policy makers, healthcare staff?

How are we facing the new changing environment?



The challenges that are facing....

- Limited support, risk assessment.....
- Lack of maintenance....
- New Biosafety Regulations"

(CWA 15793, WHO, BMBL, Canadian Standard, ISO 35001...)

- Integration of processes
- Language, cultural barriers, professional background ...





Understanding the dilemma

Substitutions-Reductions (surrogate organism) What's more difficult ...

Administrative controls *What we do not comply.....*

Work practices and procedures What we usually forget....

Engineering controls What we depend on...

Personal protective equipment (PPE) What we think is the most important....



Activities and efforts on a National level for supporting the **Epidemiological Surveillance: Collaborating Center for Laboratory Biosafety**



Impact of InDRE in the LATAM region...

 Regional experts in several topics (i.e., molecular biology, biosafety and biosecurity as well as virology).

 InDRE has 4 Collaborating Centers for WHO (Malaria, Arboviruses, Biosafety and Quality Assurance).



On a national level: CC-WHO for Lab Biosafety















The first situational diagnosis... 2014



Status of the Public Health Laboratories Network in 2013

70%	 Evidence of a biosafety manual, the implementation and documented in the Quality System of the lab
60%	 Biological safety datasheets elaborated (25%) Biological inventories updated
60%	 Evidence of a "biosafety responsible in the lab with formal authority Institutional Biosafety Commitee
45%	 Defined biosafety objectives and biological risk management policy (documented and communicated)
45%	 Presence of Biosafety Guidelines and its communication to the lab staff
45%	Evidence of the "risk communication" (signals and hazard identification)
45%	Biosafety inspections
20%	Risk assessment using Bio-RAM software or any other methodology
20%	Biological Risk Managemente Audits
0%	 Risk mitigation based on hazard identification Despite they apply actions and have documented SOPs

Event	Place/Date	Results obtained	
Biosafety and Biosecurity Workshop 3 courses	LESP Veracruz 2008, 2009 y 2010 CDC y EWIDS	23 LESP 10 InDRE	BIOSEGURIDAD 3
Biological Risk Management Workshop 3 courses	Mexico City May 2011; WHO/PAHO October 2011; CDC/EWIDS June 2012; InDRE	38 LESP 3 InDRE	
Infectious Substance Shipping Certification Course (IATA/PAHO)	InDRE November 2011 WHO/PAHO	15 LESP 8 InDRE	Hadwards & standards of the standards of the standards o
Biosafety and Biosecurity Basics in the Laboratory	InDRE November 2011 WHO/PAHO	15 LESP 8 InDRE	
<i>Other training:</i> Maintenance of Laboratory Equipment	InDRE, February 2011	31 LESP, Mainten responsibles	ance
		DRE	

The result... 2016



Current Status of the Public Health Laboratories Network

90%	 Evidence of a biosafety manual, the implementation and documented in the Quality System of the lab
90%	 Biological safety datasheets Biological inventories updated
90%	 Evidence of a "biosafety responsible in the lab with formal authority Institutional Biosafety Commitee
90%	 Defined biosafety objectives and biological risk management policy (documented and communicated)
90%	Presence of Biosafety Guidelines and its communication to the lab staff
90%	• Evidence of the "risk communication" (signals and hazard identification)
75%	Biosafety inspections
85%	Risk assessment using Bio-RAM software or any other methodology
100%	Biological Risk Managemente Audits included in the Q.S. inspections
90%	Risk mitigation based on hazard identification

Facing the challenge of new emerging diseases: National Strategy for Detecting Non-circulating Lyssaviruses in the National Public Health Laboratory Network (RNLSP)



Rabies



- In North America, rabies persists in several terrestrial meso-carnivore species and bats.
- Specific variants of the rabies virus are adapted to species as well as specific geographic areas.
- The wildlife species most commonly confirmed with rabies include skunks, foxes, coyotes, and bats.

Rabies transmission by vampire bats (*Desmodus rotundus*) is an important Public Health and economic concern in Mexico and Latin America



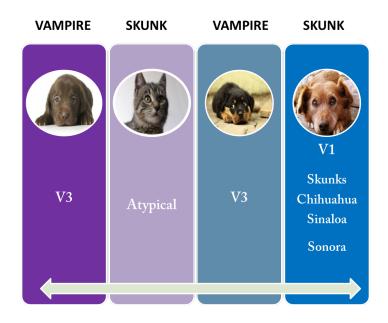
Rabies virus in Mexico

- No human cases transmitted by dogs since 2006...
- Last canine case was in 2017...

In 2017:

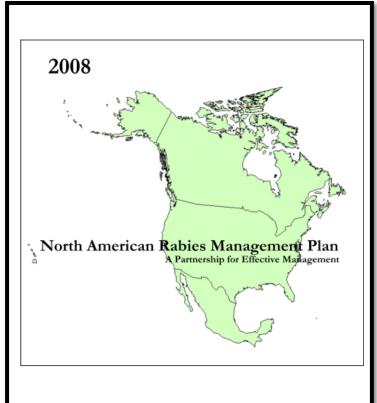
- *75% of the positive cases diagnosed were puppies or kitten....
- Confirmed cases were non-vaccinated animals...
- *All cases were related with wild rabies virus cycle





The North American Rabies Management Plan (NARMP)

- NARMP establishes a protocol for rabies management in North America by assessing and defining the needs, priorities, and strategies.
- This represents a key strategy in facilitating a planning processes by which mutual border rabies control and prevention goals and objectives can be identified and better met among Canada, Mexico, and the United States.





The North American Rabies Management Plan (NARMP)

Firmas

Plan de Control de la Rabia en Norte América

La prevendón y control de la tabla en Notre Anteista es un treto moy importente. La tabla en una anformendal virta, apuda y facial de las maniferra que contenientes en transmitida por la mortedura de asimales nichosco, los que tentidose en impessora es la sada) edidica, la agricoltare y la vida silvente. El control de la tabla significa para los gobiernos y la gente de los puises de Norteamética un costo de ciencas de militores de aditores colas alos.

nents y objetivis de prevención y control de la máis en las fronizas comunos e identificar la migra forma de alcananda por la colaboración nem Canada, Metarios y las tituados uticios. El Plan fan del techicio para disponse de ma gián comeio y parter como canadande e la las acciosas de colaboración para el control de la máis a nivel continental. Los componentes deres de teste plan ladopen comunicaciones retristrais sobre las políticas de control y la instados de la máis, miseramido en información deristrica y científia y colhadireción en processa de exigladace a propetens de control de la nais en sia frontense connance de los trate países. El objetivo suítimo dal plán, es heñadar se mace o de referencia y un dors para la interneción constructiva de los estados, paroinde ja vielendi facerdande de Canada, Meleco y las limidados da laba para enfertarser lem de forma conjunto y mediame ente plan, entre mans ageros de que las menas a lango plano para el construi de la máis en suás país señas cionandas indevidamismente y conjuntos en tada i regidi de Namaz-América.	
CANADA	
Agencia Canadiense de-Inspección de Alimentos (CPIA) Fecha	L
Arten A- Kennel Acting IV. 7/ act /200	
MEXICO (W) - 200 (3/10/03	
Secretaría de Medio Ambiente y Recursos Naturales(SEMARNAT) Fecha	L
Dirección General de Vida Silvestre	
Secretuerá de Agriculturta, Ganador Desarrollo Rural, Pesca y Alimentación (SAGARPA) Fecha Secréticio Nacional de Sanidad, Inocuidad y Calelad Agroalimentaria	
Secretaria de Saluto Epidemiológica y Control de Enfermedades (CENAVECE) Centro Netional de Vigilancia Epidemiológica y Control de Enfermedades (CENAVECE)	
LA NACIÓN NAVAJO	L
a Kugher Narajo Becha	
ESTADOS UNIDOS Servido ve subid en Humanon (HHS) Cembro de Provención y Construi de Enfermedades (CDC) Pecha	
Departamento de Agelegadora (USDA) Servicio de Inspección Santania de Planas y Animales (APHIS)	

Plan goal:

Provide a framework and forum for constructive interaction among the states and provinces and federal levels of CAN, U.S.A., and MEX to address challenges jointly, and ensure that long-term rabies management goals are met.



The North American Rabies Management Plan (NARMP)





Taxonomy of Lyssaviruses

Google Custom Search		
Experiences to show 1 Order		
Home Information Taxonomy Files Discussions Study Groups Meetings ICTV Report Login/Join		
- Realm: <i>Ribovine</i> 1 phylu	m. 3 orders. 40 families. 8 genera	
- Phylum: Negarnaviricota Member of Riboviria	2 subphyla	history
Subphylum: Haploviricotina Member of Negarnaviricota	4 classes	<u>history</u>
- Class: Chunqiuviricetes Member of Haploviricotina	1 order	<u>history</u>
+ Order: Muvirales Member of Chunqiuviricetes	1 family	<u>history</u>
Class: Milneviricetes Member of Haploviricotina	1 order	<u>history</u>
+ Order: Serpentovirales Member of Milneviricetes	1 family	<u>history</u>
Class: Monjiviricetes Member of Haploviricotina	2 orders	<u>history</u>
+ Order: Jingchuvirales Member of Monjiviricetes	1 family	<u>history</u>
	11 families	<u>history</u>
+ Family: Artoviridae Member of Mononegavirales	1 genus	history
+ Family: Bornaviridae Member of Mononegavirales	3 genera	history
+ Family: Filoviridae Member of Mononegavirales	5 genera	history
+ Family: Lispiviridae Member of Mononegavirales	1 genus	history
+ Family: Mymonaviridae Member of Mononegavirales	1 genus	history
+ Family: Nyamiviridae Member of Mononegavirales	6 genera	history
+ Family: Paramyxoviridae Member of Mononegavirales	4 subfamilies, 3 species	history
+ Family: Pneumoviridae Member of Mononegavirales	2 genera	history
+ Family: Rhabdoviridae Member of Mononegavirales	20 genera, 1 species	history
+ Family: Sunviridae Member of Mononegavirales	1 genus	history
+ Family: Xinmoviridae Member of Mononegavirales	1 genus	history
Class: Yunchangviricetes Member of Haploviricotina	1 order	history
+ Order: Goujianvirales Member of Yunchangviricetes	1 family	history
- Subphylum: Polyploviricotina Member of Negarnaviricota	2 classes	history
- Class: Ellioviricetes Member of Polyploviricotina	1 order	history

Source: ICTV Mayo 2019 https://talk.ictvonline.org/taxonomy/



Taxonomy of Lyssaviruses (cont.)



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Information

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Taxonomy rices Discussions Study Groups meetings for viteport Loginoon		
Family: Paramyxoviridae Member of Mononegavirales	4 subfamilies, 3 species	history
Family: Pneumoviridae Member of Mononegavirales	2 genera	history
- Family: Rhabdoviridae Member of Mononegavirales	20 genera, 1 species	history
+ Genus: Almendravirus Member of Rhabdoviridae	5 species	
+ Genus: Alphanemrhavirus Member of Rhabdoviridae	2 species	<u>history</u>
+ Genus: Caligrhavirus Member of Rhabdoviridae	3 species	
+ Genus: Curiovirus Member of Rhabdoviridae	4 species	<u>history</u>
+ Genus: Cytorhabdovirus Member of Rhabdoviridae	11 species	
+ Genus: Dichorhavirus Member of Rhabdoviridae	5 species	
+ Genus: Ephemerovirus Member of Rhabdoviridae	8 species	
+ Genus: Hapavirus Member of Rhabdoviridae	15 species	<u>history</u>
+ Genus: Ledantevirus Member of Rhabdoviridae	16 species	
+ Genus: Lyssavirus Member of Rhabdoviridae	16 species	<u>history</u>
+ Genus: Novirhabdovirus Member of Rhabdoviridae	4 species	<u>history</u>
+ Genus: Nucleorhabdovirus Member of Rhabdoviridae	10 species	
+ Genus: Perhabdovirus Member of Rhabdoviridae	3 species	
+ Genus: Sigmavirus Member of Rhabdoviridae	7 species	<u>history</u>
+ Genus: Sprivivirus Member of Rhabdoviridae	2 species	<u>history</u>
+ Genus: Sripuvirus Member of Rhabdoviridae	5 species	<u>history</u>
+ Genus: Tibrovirus Member of Rhabdoviridae	7 species	
+ Genus: Tupavirus Member of Rhabdoviridae	3 species	<u>history</u>
+ Genus: Varicosavirus Member of Rhabdoviridae	1 species	<u>history</u>
+ Genus: Vesiculovirus Member of Rhabdoviridae	16 species	
Species: Moussa virus Member of Rhabdoviridae		<u>history</u>

1



New species submitted

Source: ICTV Mayo 2019 https://talk.ictvonline.org/taxonomy/

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ICTV Taxonomy history: Rabies lyssavirus

Riboviria > Negamaviricota > Haploviricotina > Monjiviricetes > Mononegavirales > Rhabdoviridae > Lyssavirus > Rabies lyssavirus

2018b EC 50, Washington, DC, July 2018; Email ratification February 2019 (MSL #34)

Moved

Riboviria > Negarnaviricota > Haploviricotina > Monjiviricetes > Mononegavirales > Rhabdoviridae > Lyssavirus > Rabies lyssavirus

Proposal

2018a EC 50, Washington, DC, July 2018; Email ratification October 2018 (MSL #33)

Moved

Negamaviricota > Haploviricotina > Monjiviricetes > Mononegavirales > Rhabdoviridae > Lyssavirus > Rabies lyssavirus

Proposal

2015 EC 47, London, UK, July 2015; Email ratification 2016 (MSL #30)

Renamed

Mononegavirales > Rhabdoviridae > Lyssavirus > Rabies lyssavirus

Proposal

1990 Plenary session vote 29 August 1990 in Berlin (MSL #11)

Moved

Mononegavirales > Rhabdoviridae > Lyssavirus > Rabies virus

Proposal

1975 Plenary session vote 12/16 September 1975 in Madrid (MSL #03)

Moved, Assigned as Type Species Rhabdoviridae > Lyssavirus > Rabies virus

Proposal

1971 ICTV 1st Report (MSL #01)

New

Rhabdovirus > Rabies virus

Proposal

Evolution in the classification of Lyssaviruses

SEROTYPE (1970)

Antibody cross reaction : Rabies, Lagos bat virus, Mokola virus, Duvenhage virus

PANEL OF MONOCLONAL ANTIBODIES (1978)

Rabies virus antigenic variants in order to determine the most probable reservoir species

SEROGENOTYPE (1980)

Rabies, Lagos bat virus , Mokola virus , Duvenhage virus, EBLV-1, EBLV-2, Australian bat lyssavirus.

SPECIES (2003)

16 identified species officially (2019)



Genus Lyssavirus

Virus Taxonomy: 2018b Release

EC 50, Washington, DC, July 2018 Email ratification February 2019 (MSL #34)

— Genu	s: Lyssavirus
Species:	Aravan lyssavirus
Species:	Australian bat lyssavirus
Species:	Bokeloh bat lyssavirus
Species:	Duvenhage lyssavirus
Species:	European bat 1 lyssavirus
Species:	European bat 2 lyssavirus
Species:	Gannoruwa bat lyssavirus
Species:	Ikoma lyssavirus
Species:	Irkut lyssavirus
Species:	Khujand lyssavirus
Species:	Lagos bat lyssavirus
Species:	Lleida bat lyssavirus
Species:	Mokola lyssavirus
Species:	Rabies lyssavirus
Species:	Shimoni bat lyssavirus

Species: West Caucasian bat lyssavirus

Derivation of names

Lyssavirus: from *Lyssa*, the Greek goddess of madness, rage, and frenzy.

Related, unclassified viruses

Virus name	Accession number	Virus abbreviation
Taiwan bat lyssavirus	MF472710	TBLV
Kotalahti bat lyssavirus	MF960865	KBLV

Virus names and virus abbreviations are not official ICTV designations.

Source: ICTV Mayo 2019 https://talk.ictvonline.org/taxonomy/



Genomic Organization of the Lyssaviruses

rabies virus 3' - N Australian bat lyssavirus 3' N Duvenhage virus European bat lyssavirus 1 3' € European bat lyssavirus 2 3' L N Aravan virus 3' N Khujand virus 3' N Bokeloh bat lyssavirus 3' [N Irkut virus 3' N Gannoruwa bat lyssavirus 3' N Lagos bat virus 3' N Mokola virus 3' N Shimoni bat virus 3' N West Caucasian bat lyssavirus 3' Ikoma virus Lleida bat lyssavirus

- N, P, M, G and L represent the ORFs that code for structural proteins.
- Genomes have untranslated regions in the *G* genes; In the West Caucasus Bat Lyssavirus (WCBL), this region ~ 700 nt contains an ORF of 180nt (in blue) that encodes a possible 7.1 kDa protein.
- The West Caucasus Bat Lyssavirus also has an alternative ORF of 228 nt (orange), which begins near the beginning of the M gene, which encodes a putative protein of 8.4 kDa. It is not known if these proteins are expressed in infected cells.

Source: ICTV Mayo 2019: <u>https://talk.ictvonline.org/ictv-reports/ictv_online_report/negative-sense-rna-viruses/mononegavirales/w/rhabdoviridae/795/genus-lyssavirus</u>



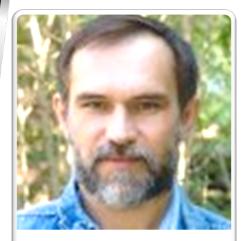
13 kb

- Phylogroup I

Phylogroup II

New species

Rabies virus (RABV) (1865) Lagos bat lyssavirus (LBV) (1956) Mokola virus(MOKV) (1974) Duvenhage virus (DUVV) (1971) European bat lyssavirus 1 [EBLV-1] (1968) European bat lyssavirus 2 [EBLV-2] (1986) Australian bat lyssavirus (ABLV) (1996) Aravan virus (ARAV) (2003) lrkut virus (IRKV) (2003) Khujand virus (KHUV) (2003) West Caucasian Bat Virus (WCBV) (2005) Shimoni virus (SHIBV (2009) Bokeloh bat lyssavirus (BBLV) (2010) lkoma lyssavirus (IKOV) (2012) Lleida bat lyssavirus (LLEBV) (2012) Gannoruwa bat lyssavirus (GBLV) (2016) Taiwan bat lyssavirus (TWBLV) (2016/2017) Kotalahti bat lyssavirus (KBLV) (2017)



Dr. Iván Kuzmin

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PHYLOGROUPS

Virus Distribution No. RABV Global * 1 4 3 ARAV 2 Eurasian 🛶 KHUV Eurasian 🛶 3 BBLV European 🛩 4 EBLV-2 Europe 🔸 5 6 ABLV Australia 🛶 IRKV Eurasian 🛶 EBLV-1 European 🛶 8 DUVV African 🔸 9 Phylogroup I Phylogroup II 0.1 African MOKV 10 SHIBV 11 African + LBV 10 12 African * 12 Ŕ Ď Phylogroup III/IV? 13 WCBV Eurasian 🛶 13 14 IKOV African 14 15 15 LLEBV European 🔸

Source: Lyssaviruses and Bats: Emergence and Zoonotic Threat. Viruses 2014, 6, 2974-2990; doi:10.3390/v6082974



Relationship between Lyssaviruses and bats species

Geographical distribution	Lyssavirus species	Bat species most commonly associated with lyssavirus infection	Common name	Transmission from bats implicated in human fatalities
The Americas	Rabies virus (RABV)	Eptesicus fuscus	Big brown bat	Yes
		Tadarida brasiliensis	Mexican/Brazilian free-tail bat	Yes
		Lasionycteris noctivagens	Silver-haired bat	Yes
		Perimyotis subflavus	Tri-coloured bat	Yes
		Desmodus rotundus	Vampire bat	Yes
Africa	Lagos Bat Virus (LBV)	Eidolon helvum	Straw coluored fruit bat	No
		Rousettus aegyptiacus	Egyptian fruit bat	No
		Epomorphorus wahlbergi	Wahlberg's epauletted fruit bat	No
	Shimoni Bat Virus (SHIBV)	Hipposideros commersoni	Commerson's leaf-nosed bat	No
	Duvenhage virus (DUVV)	Miniopterus sp?	Undefined	Yes
		Nycteris thebaica	Egyptian slit-faced bat	Yes
Eurasia	European Bat Lyssavirus type 1 (EBLV-1)	Eptesicus serotinus	Serotine bat	Yes
	European Bat Lyssavirus type 2 (EBLV-2)	Myotis daubentonii	Daubenton's bat	Yes
	Bokeloh Bat Lyssavirus (BBLV)	Myotis nattereri	Natterer's bat	No
	Aravan virus (ARAV)	Myotis blythi	Lesser mouse-eared bat	No
	Irkut Virus (IRKV)	Murina leucogaster	Greater tube-nosed bat	Yes
	Khujand Virus (KHUV)	Myotis mystacinus	Whiskered bat	No
	West Caucasian Bat Virus (WCBV)	Miniopterus schreibersii	Common bent-winged bat	No
	Lleida Bat Lyssavirus (LLEBV) *	Miniopterus schreibersii	Common bent-winged bat	No
Australasia	Australian Bat Lyssavirus (ABLV)	Pteropus alecto	Black flying fox and related sp.	Yes
		Saccolaimus flaviventris	Yellow-bellied sheath-tailed bat	Yes



* Genetic data only reported to LLEBV

However.....

What to do when a non endemic *Lyssavirus* can be potentially introduced into a different region?







Actions implemented:

A strategic measure for increasing Rabies and other
 Lyssaviruses surveillance in the country and action plan for
 the containment of these pathogens was developed in the
 country.....



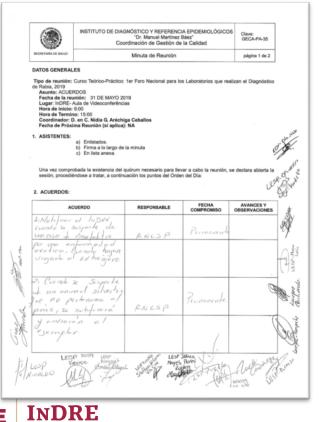
Purposes of the action plan

- Provide timely response.....
- Guarantee the development of activities under the highest standards of quality and safety.
- Provide advice and support to specific response groups....



Building the Future: Agreement with the Rabies Lab Network 2018-2019

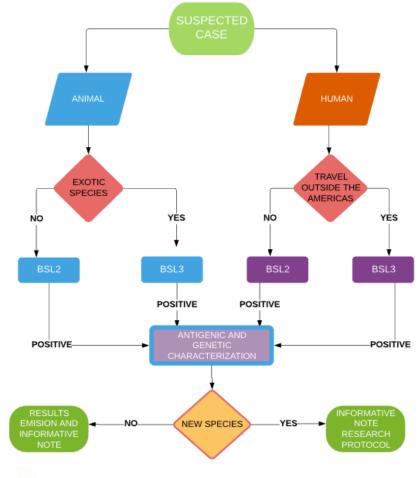






What to do against a noncirculating Lyssavirus in the region?







Importance of the containment lab (CL) for the epidemiological surveillance

- It has a unique and innovative design for the safe and secure handling of high-risk pathogens....
- It has a defined installed capacity for a large Public Health emergency response....
 - "Adaptability of the facilities" that can be used for diagnostic/research purposes.



Diagnosis at the CL

ER based on:

- 1.- Activities, frequency, complexity and time required.....
- 2.- Special requirements (i.e., vaccination).
- 3.- Special situations (i.e., epidemiological emergencies).

 Table 1. Risk assessment for the use of containment facilities at InDRE by lab

Laboratory	Status			
Viral pathogens				
Arboviruses and HF				
Rabies				
Poliovirus				
Respiratory viruses				
Febrile Exhantematic viruses				
Special pathogens				

High Priority 🦰 Medium Priority 📘 Low Priority



What have we been doing in terms of biosafety and biosecurity for support the PH labs?

- Frequent training from 2008 to date....
- Specific emergency response drills for possible scenarios evolving containment lab.....
- More than 120 trained staff for working at the CL.....
- Audits, inspections and risk communication....











Additional measures

- National cooperation with other agencies....
- Strategic communication with international reference centers....
- Trained and certified shippers for the safe transport of Infectious Substances





Concluding remarks (International Level)

What we need to do:

- Improve the exchange of information on epidemiologic events...
- Promote collaborative responses...
 - Facilitate the international cooperation



Concluding remarks (National Level)

- Successful transfer of technology
- Specific protocols based on WHO recommendations and other specialized centers (i.e., CDC, PHAC).



Concluding remarks (cont.)

- More than 25 guidelines for different diseases in the country.
- Emergency preparedness and response protocols (i.e., Ebola outbreak National Response Plan since 2014).
- Strong Quality Assurance processes (ISO 9001:2015, ISO 15189:2012 as well as, CWA 15793 and CWA 16335 standards).
- Experience and training (basic and specific).





Success depends on a TEAM.....



General Direction of Epidemiology

Direction of Diagnosis and Reference Direction of Services and Technical Support

Department of Virology Department of Bacteriology Department of Parasitology Department of Molecular Biology Department of Emerging Diseases Department of Sample Control and Services

RNLSP



All is about emerging and re-emerging infectious diseases/pathogens.....

"Sooner or later everything old is new again".....



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Thank you for your attention

Merci de votre attention

