

Partnerships and Collaborations to advance the
vaccine agenda:
Lessons from the Rotavirus Experience

Roger I. Glass, M.D., Ph.D.
Fogarty International Center, NIH
Viral Gastroenteritis Unit, CDC

IOM Estimates of Rotavirus Deaths--1985

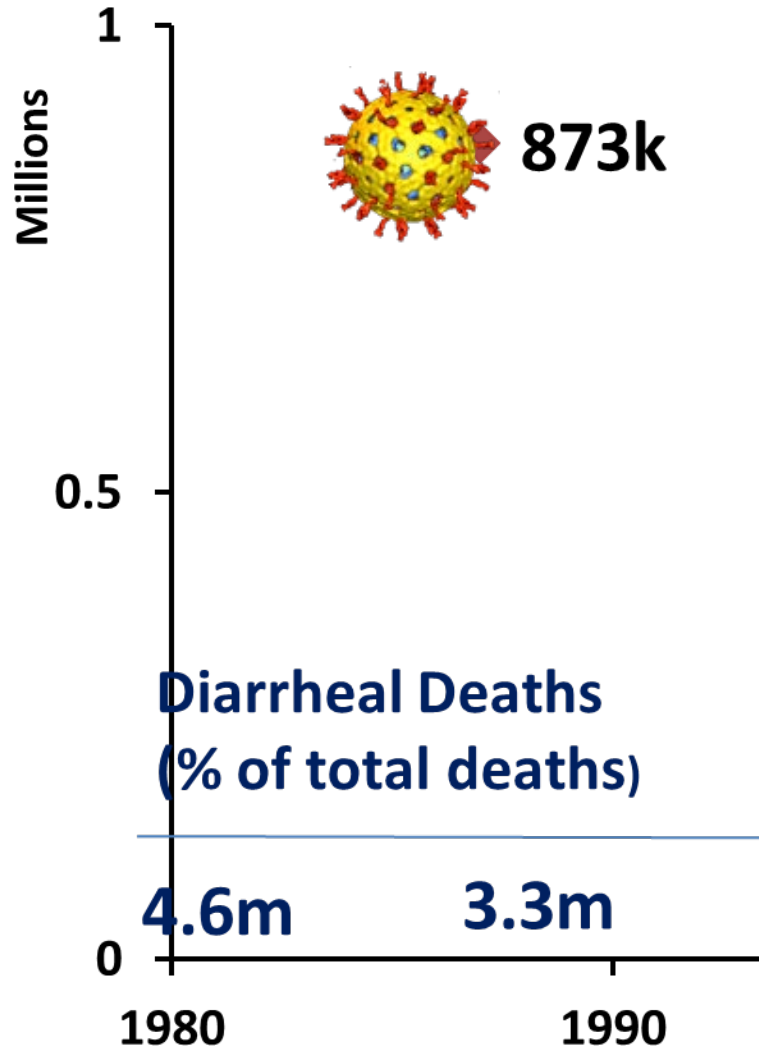
 INSTITUTE OF MEDICINE
OF THE NATIONAL ACADEMIES

New Vaccine Development Establishing Priorities

VOLUME II

Diseases of Importance in Developing Countries

Part Two of a Two-Part Study by the Committee on
Issues and Priorities for New Vaccine Development

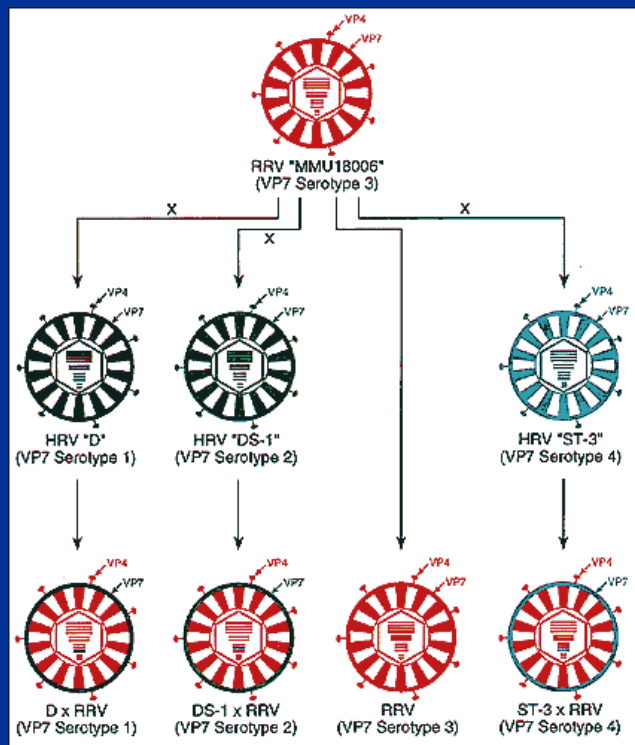


***High priority for
Developing
Countries...***

1998 Rhesus Tetraivalent vaccine-Rotashield Licensed



Albert Kapikian,



INFECTIOUS DISEASES IN CHILDREN
Rotavirus vaccine shown to be cost effective
Oberlin, Ohio
February 1997

NEWS VS. HISTORY
Honor due to Kapikian
Washington - When Washington State's first and only...
Winchester-Salem Journal
September 8, 1995

The Virginian-Pilot
New vaccine may tame common childhood virus

The New York Times
F.D.A. Approves Vaccine for Childhood Diarrhea
By THE ASSOCIATED PRESS
Washington - The Food and Drug Administration Monday approved the first vaccine against a leading cause of childhood diarrhea, a virus that hospitalizes 55,000 American children a year and kills one million in other countries.

HEALTHCARE REVIEW
Southern New England Edition (Massachusetts, Connecticut & Rhode Island)
Circulation: 52,000
Date: Feb./Mar./March 1997
Rotavirus: Affecting children and healthcare costs across the globe

Vaccine offers way to prevent child diarrhea

Pediatric News
Circulation: 15,416
Date: November 1996
Rotavirus Vaccine Cuts Diarrhea Hospitalizations



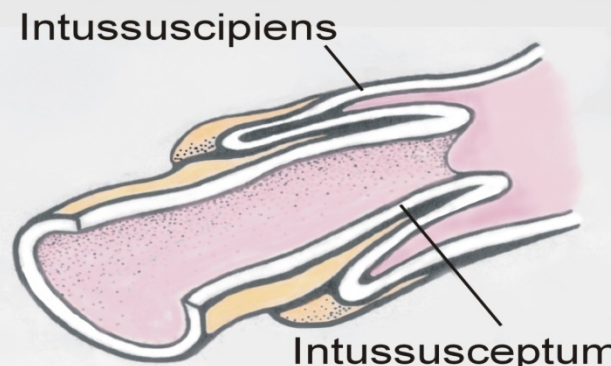
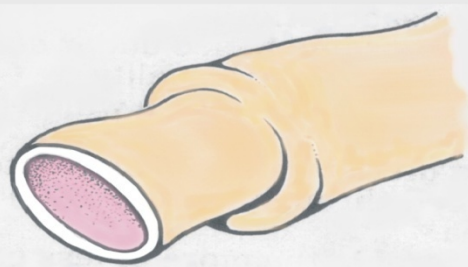
MMWR™

MORBIDITY AND MORTALITY
WEEKLY REPORT

- 577 Intussusception Among Recipients of Rotavirus Vaccine — United States, 1998–1999
- 582 Outbreak of *Salmonella* Serotype Muenchen Infections Associated with Unpasteurized Orange Juice — United States and Canada, June 1999
- 585 Progress Toward Measles Elimination — Southern Africa, 1996–1998
- 590 Recommendations of the Advisory Committee on Immunization Practices: Revised Recommendations for Routine Poliomyelitis Vaccination

Intussusception Among Recipients of Rotavirus Vaccine — United States, 1998–1999

On August 31, 1998, a tetravalent rhesus-based rotavirus vaccine (RotaShield®*, Wyeth Laboratories, Inc., Marietta, Pennsylvania) (RRV-TV) was licensed in the United States for vaccination of infants. The Advisory Committee on Immunization Practices



Intussusception

The telescoping of the intestine onto itself usually at the ileal-cecal junction, leading to reversible repair or entrapment with edema, necrosis and perforation



1990s

**DEPARTMENT OF
MEDICAL RESEARCH**



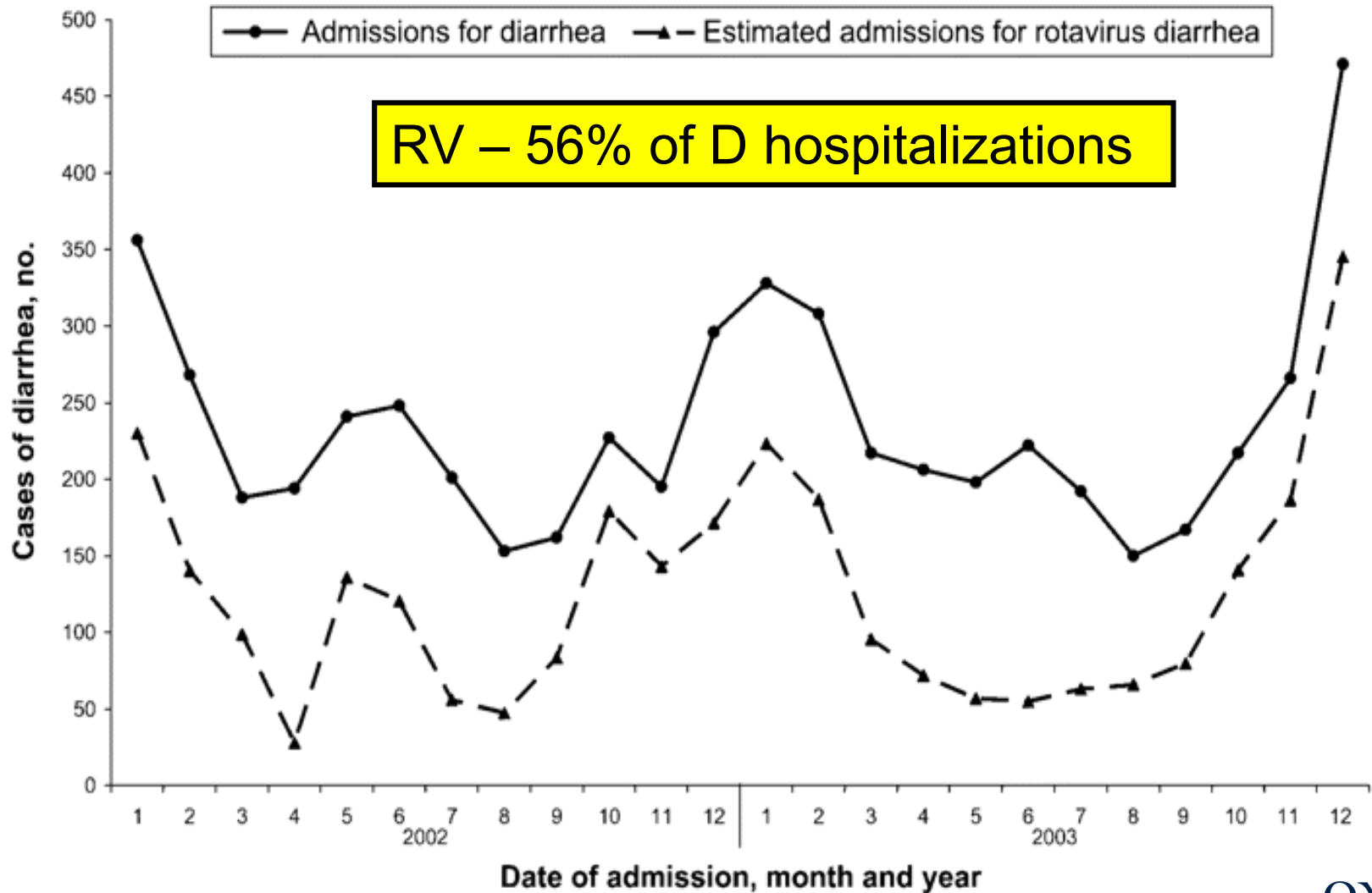
Kyaw Moe
Virology Research Div.

Hospital-Based Surveillance for Rotavirus Diarrhea in Children in Yangon, Myanmar

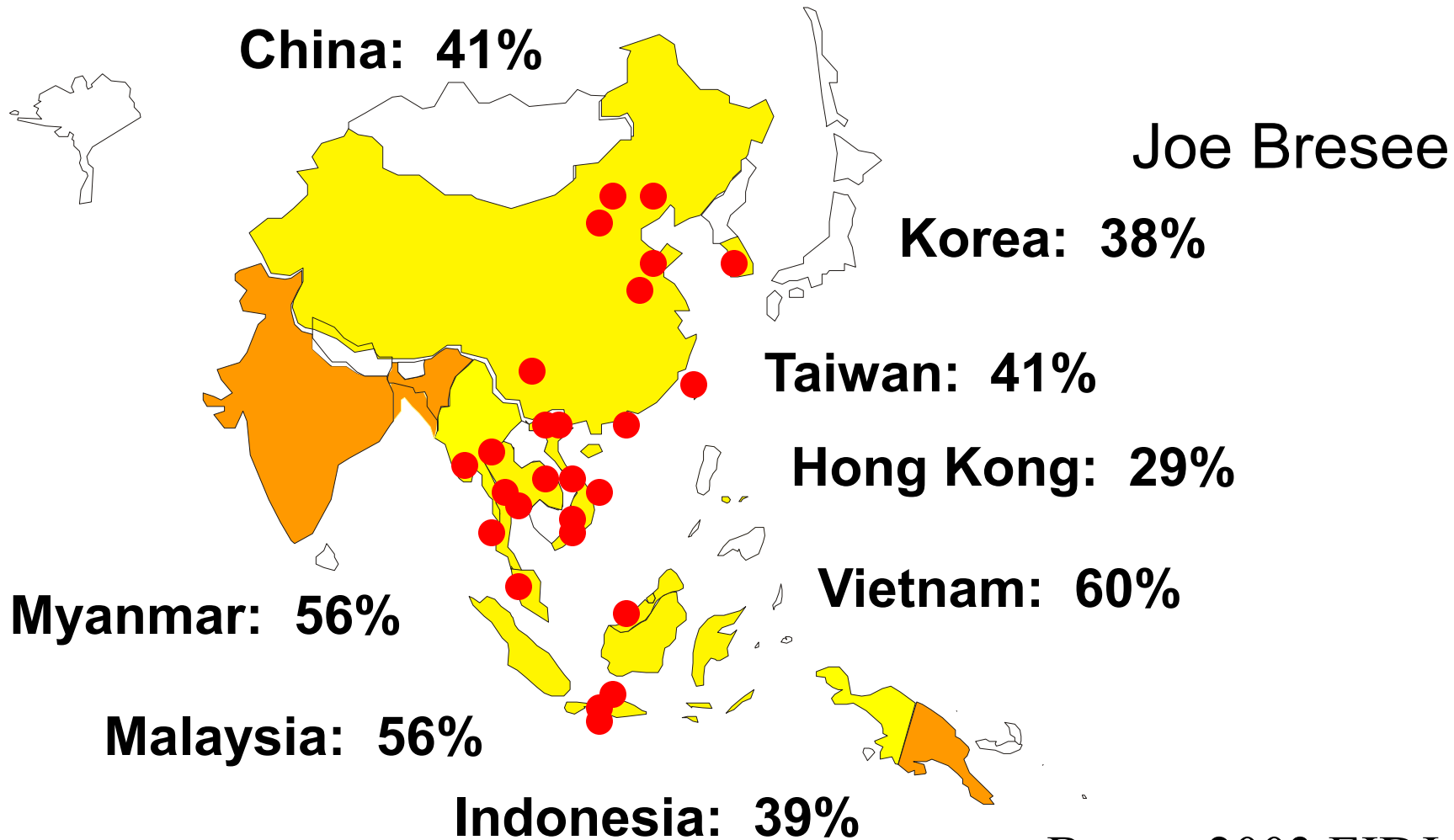
Kyaw Moe,¹ Erik G. Hummelman,² Win Mar Oo,¹ Thandar Lwin,¹ and Tin Tin Htwe¹

¹Virology Research Division, Department of Medical Research (Lower Myanmar), Yangon, Myanmar; ²Centers for Disease Control and Prevention, Atlanta, Georgia

Diarrhea-associated admissions of children <5 years to Yangon Children's Hospital , January 2002 - December 2003.



Rotavirus Hospitalizations in the Asian Rotavirus Surveillance Network



Joe Bresee

Bresee 2003 EIDJ



Tools for Global Rotavirus Surveillance

Laboratory Manual



Sentinel Hospital Surveillance

Cost- Effectiveness

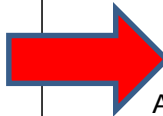
Impact Evaluation



Percent with Rotavirus (N ~58)

WPR

SEAR

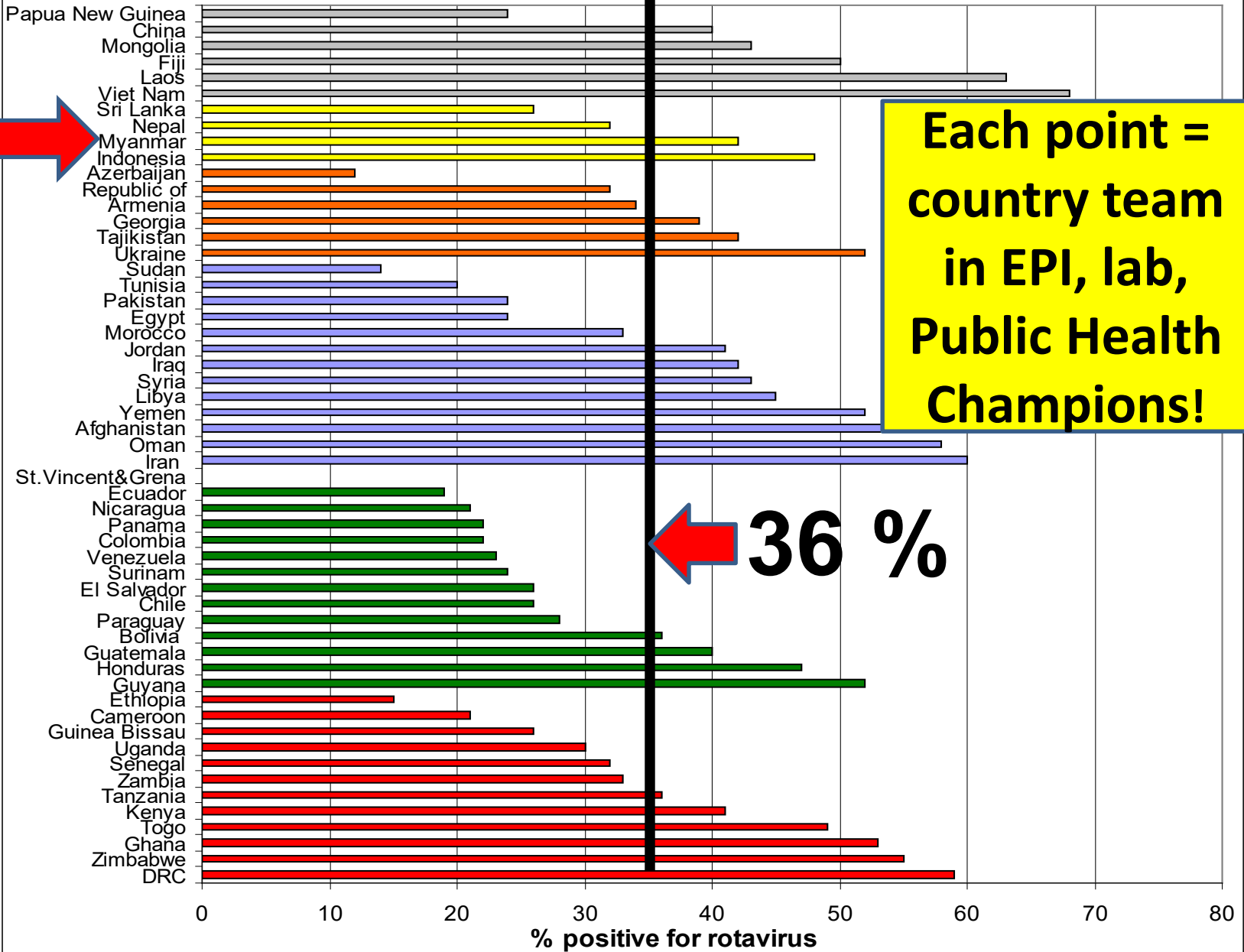


EUR

EMR

AMR

AFR



Each point = country team in EPI, lab, Public Health Champions!

36 %

% positive for rotavirus



Contents lists available at ScienceDirect

Vaccine

journal homepage: www.elsevier.com/locate/vaccine



Review

Rotavirus epidemiology: The Asian Rotavirus Surveillance Network[☆]

E.A.S. Nelson^{a,*}, J.S. Bresee^b, U.D. Parashar^b, M.-A. Widdowson^b, R.I. Glass^c,
the members of the Asian Rotavirus Surveillance Network^d

^a Department of Paediatrics, The Chinese University of Hong Kong, Hong Kong SAR, China

^b Centers for Disease Control and Prevention, Atlanta, GA, United States

^c Fogarty International Center, National Institutes of Health, Bethesda, MD, United States

^d Asian Rotavirus Surveillance Network Members—China: Zhao-Yin Fang, Bei Wang, Li-Jie Zhang, Li-Wei Sun, Zeng-Qing Du, Jing-Yu Tang, A Wang, Zhi-Yi Xu, Ying-Lin Zhang, Shou-Jun Zhao, Zhi-Yong Hao, Zhan-Chun Xing, Chang-Quan Han, Jing-Chen Ma, Ji-Chao Chen; Hong K Yu, Ying-Chu Ng, Kin-Hung Poon, Chi-Hang Ng, Kin-Sing Ip, Tai-Fai Fok; India: Rajiv Bahl, Pratima Ray, Swati Subodh, Prashant Shambh Kang, Shobhana D. Kelkar, Shoba D. Chitambar, Pratima Ray, Trailokyanath Naik. Indonesia: Yati Soenarto, Siswanto Agus Wilopo, Abu Nenny Sri Mulyani; Japan: Toyoko Nakagomi, Osamu Nakagomi, Yoshihiro Takahashi, Masamichi Enoki, Takashi Suzuki; Korea: Jung S. Sae A. Min, Tae H. Park, Dae S. Jo, Paul E. Kilgore, Batmunkh Nyambat, Zhi Y. Xu, Lorenz von Seidlein, Oak Pil Han, John Clemens; Malaysia Lailanor H. J. Ibrahim, Ahmad Faudzi H. J. Yusoff, Lee Gaik Chan; Mynamar: Kyaw Moe, Win Mar Oo, Thandar Lwin, Tin Tin Htwe; Tai Tang, Yung-Feng Huang, Ping-Ing Lee, Jyh-Yuan Yang, Hour-Young Chen; Thailand: Chuleeporn Jiraphongsa, Yaowapa Pongsuwanna, P Arporntip, Manas Kanoksil, Nakorn Prem Sri, Utcharee Intusoma; Vietnam: Nguyen Van Man, Le Thi Luan, Dang Duc Trach, Nguyen Thi H Dang Duc Anh; United States: Erik Hummelman, Jon R. Gentsch, Thea K. Fischer, Vincent P. Hsu, Ashley R. Laird, Brittany Bielfelt, Dixie Baoming Jiang, Laura Jean Podewils, Lynn Antil, Richard Rheingans, T. Christopher Mast; World Health Organization and collaborating laboratories: Bernard Hanjaj, Duncan Steele, Carl D. Kirkwood, Krisztiaín Băi nyai, Nigel A. Cunliffe



OXFORD
UNIVERSITY PRESS

Surveillance of rotavirus gastroenteritis (2015-2017); vital information for pre-and post-rotavirus vaccination in Myanmar

¹Theingi Win Myat, ¹Hlaing Myat Thu, ²Ye Myint Kyaw,
¹Nang Sarm Hom, ¹Myat Mo Zar Kham, ¹Win Mar, ¹Khin
Sandar Aye, ¹Hla Myo Thu and ¹Kyaw Zin Thant

¹Department of Medical Research

²Yangon Children Hospital



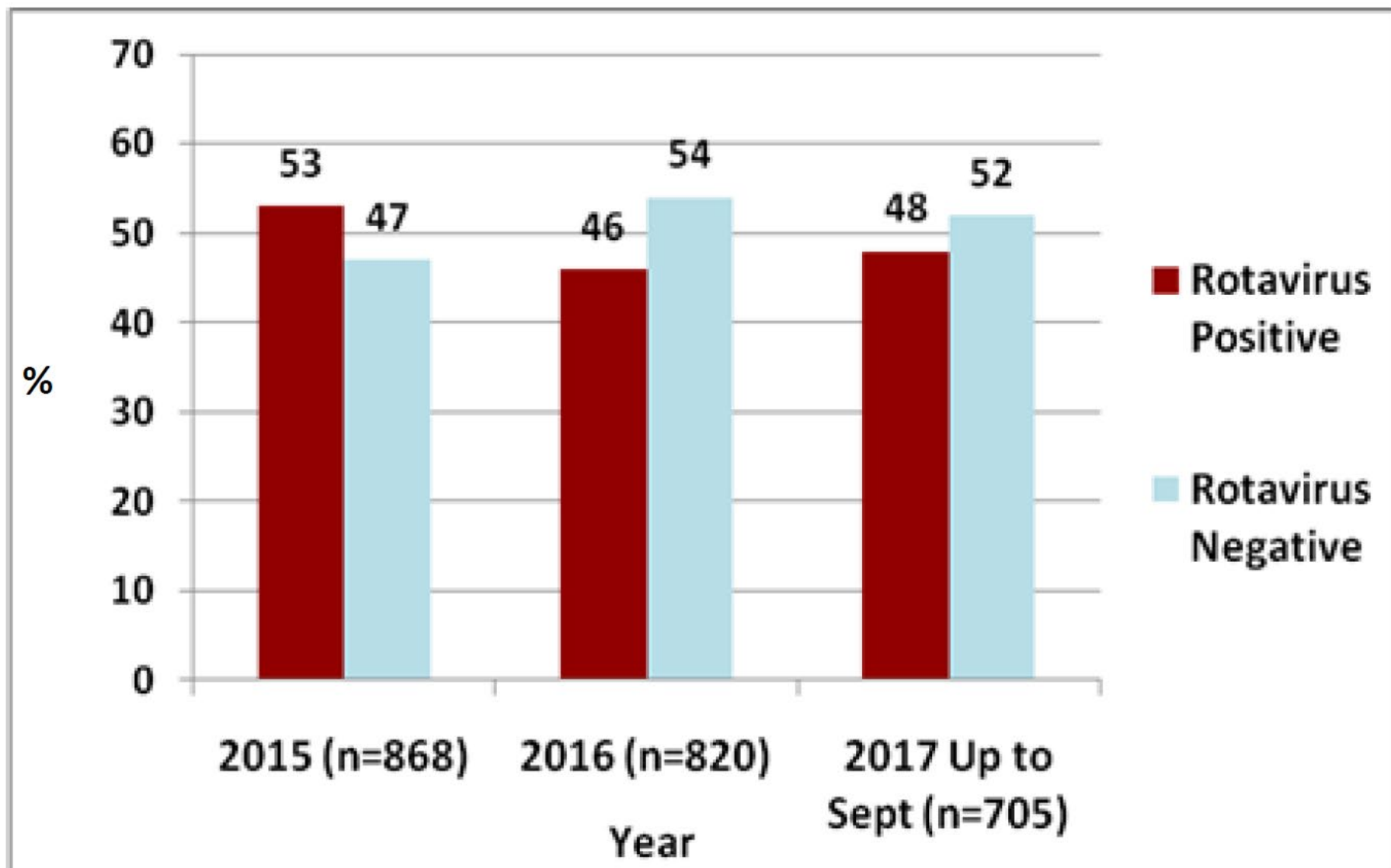


Fig (1) Proportion of Rotavirus positive cases tested by ELISA

6. Genotype distribution

Seasonal Year	Most Prevalent Genotype	N (%)	Total genotyped
2008-2009	G1P[8]	15 (35%)	43
2009-2010	G12P[8]	50 (61.7%)	81
2010-2011	G12P[8]	103(75.2%)	137
2011-2012	G12P[8]	45 (26%)	173
2012-2013	G2P[4]	22 (73.3%)	30
2013-2014	G1P[8]	31 (41.9%)	74
2014-2015	G9P[8]	72 (53.3%)	135
2015-2016	G9P[8]	25 (30.9%)	81
2016-2017	G3P[8]	32 (58.2%)	55



Inside this issue

Fogarty study shows poor-quality malaria drugs causing drug resistance p. 4

FOGARTY INTERNATIONAL CENTER • NATIONAL INSTITUTES OF HEALTH • DEPARTMENT OF HEALTH AND HUMAN SERVICES

Fogarty director joins US health delegation to Burma

As part of the warming of relations between the U.S. and Burma, Fogarty Director Dr. Roger I. Glass recently traveled to Rangoon to represent the U.S. at an international science meeting and to discuss possible research collaborations with the country's health minister.

"The Burmese are very enthusiastic about engaging with the U.S. scientific community to advance their research efforts in HIV/AIDS, malaria, dengue, and maternal and child health in particular," observed Glass. "I was very impressed with the caliber of scientists I met and look forward to establishing research and training collaborations with them."

In his role as lead U.S. health representative, Glass participated in the meeting of ASEAN's Committee on Science and Technology. He proposed that Fogarty and its federal partners host an influenza workshop in the region later this year, which was met with enthusiasm.

Courtesy of U.S. State Department



**Prof. Pe Thet Khin,
Minister of Health
Pediatrician**

Fogarty Director Dr. Roger I. Glass visited Burma recently to discuss possible research collaborations with the country's health minister, Dr. Pe Thet Khin.

Government set to introduce vaccine for rotavirus

MYINT KAY THI | 05 JUN 2015



The Ministry of Health plans to launch an immunisation drive against the deadly rotavirus in 2017-18, an official said last week.

MYANMARTIMES

Vaccine alliance gives \$60 million for immunisation programme

Mar 6, 2018

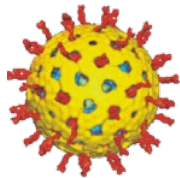
MYINT KAY THI | 06 MAR 2018



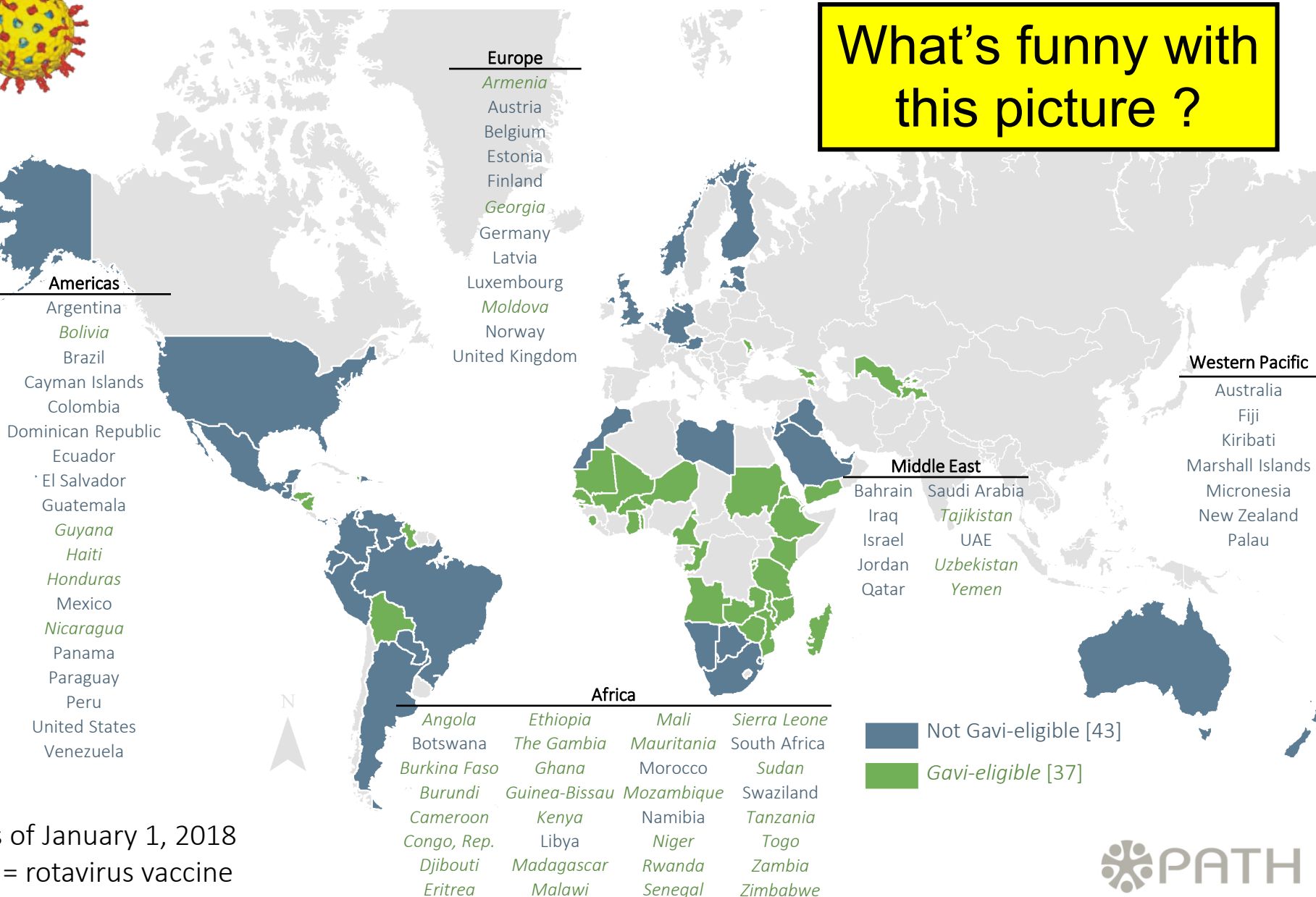
Geneva-based **Gavi, The Vaccine Alliance**, will provide **US\$60 million** over the next two years to implement an immunisation programme in the country, largely focusing on cold chain facilities, according to the Ministry of Health and Sports.

<https://www.mmtimes.com/news/vaccine-alliance-gives-60-million-immunisation-programme.html>

National RV introductions by region: 92 countries*

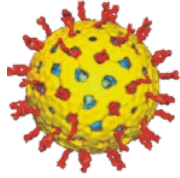


What's funny with this picture ?



*As of January 1, 2018
RV = rotavirus vaccine

National RV introductions by geographic region:



What's happened in Asia ?

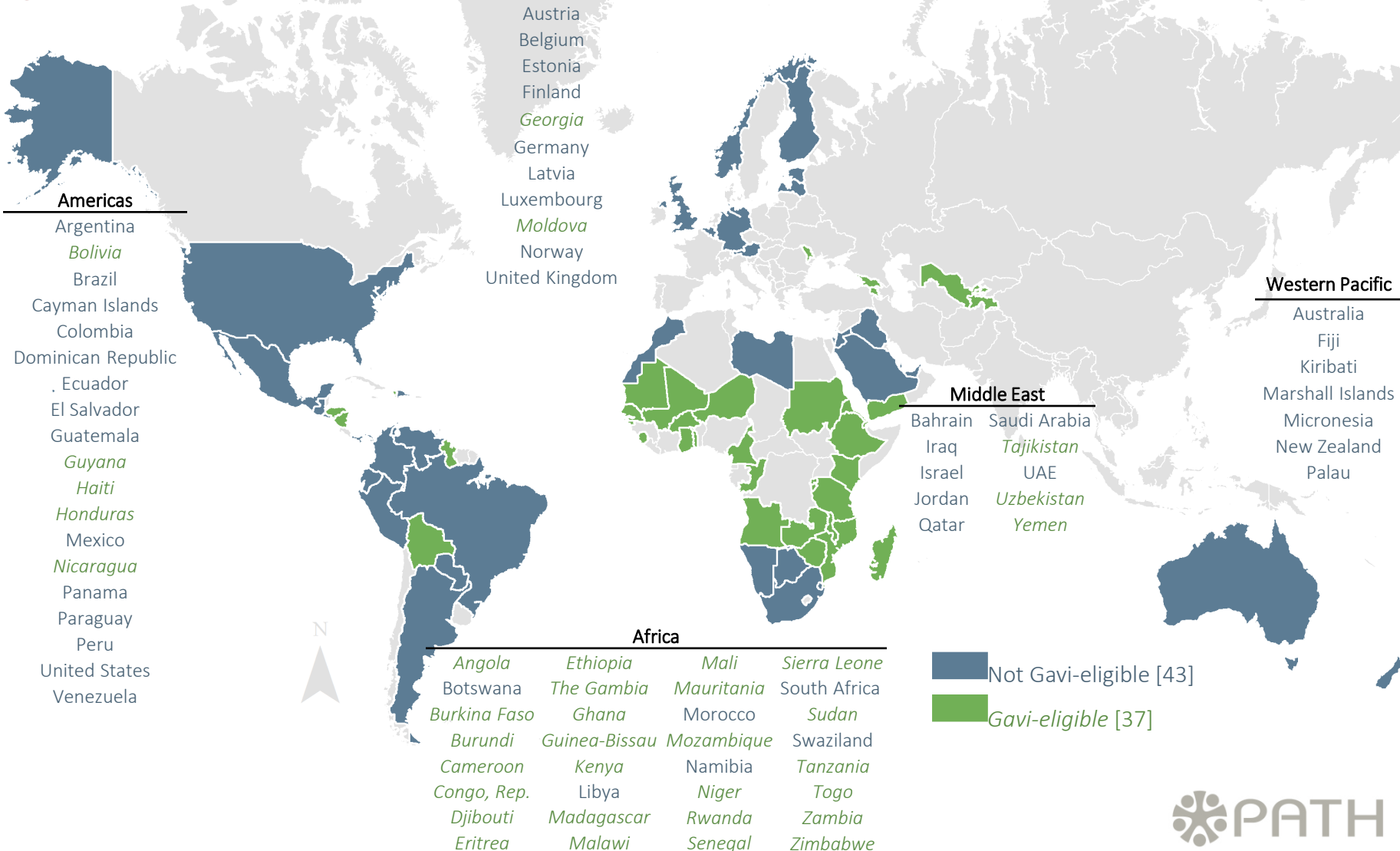
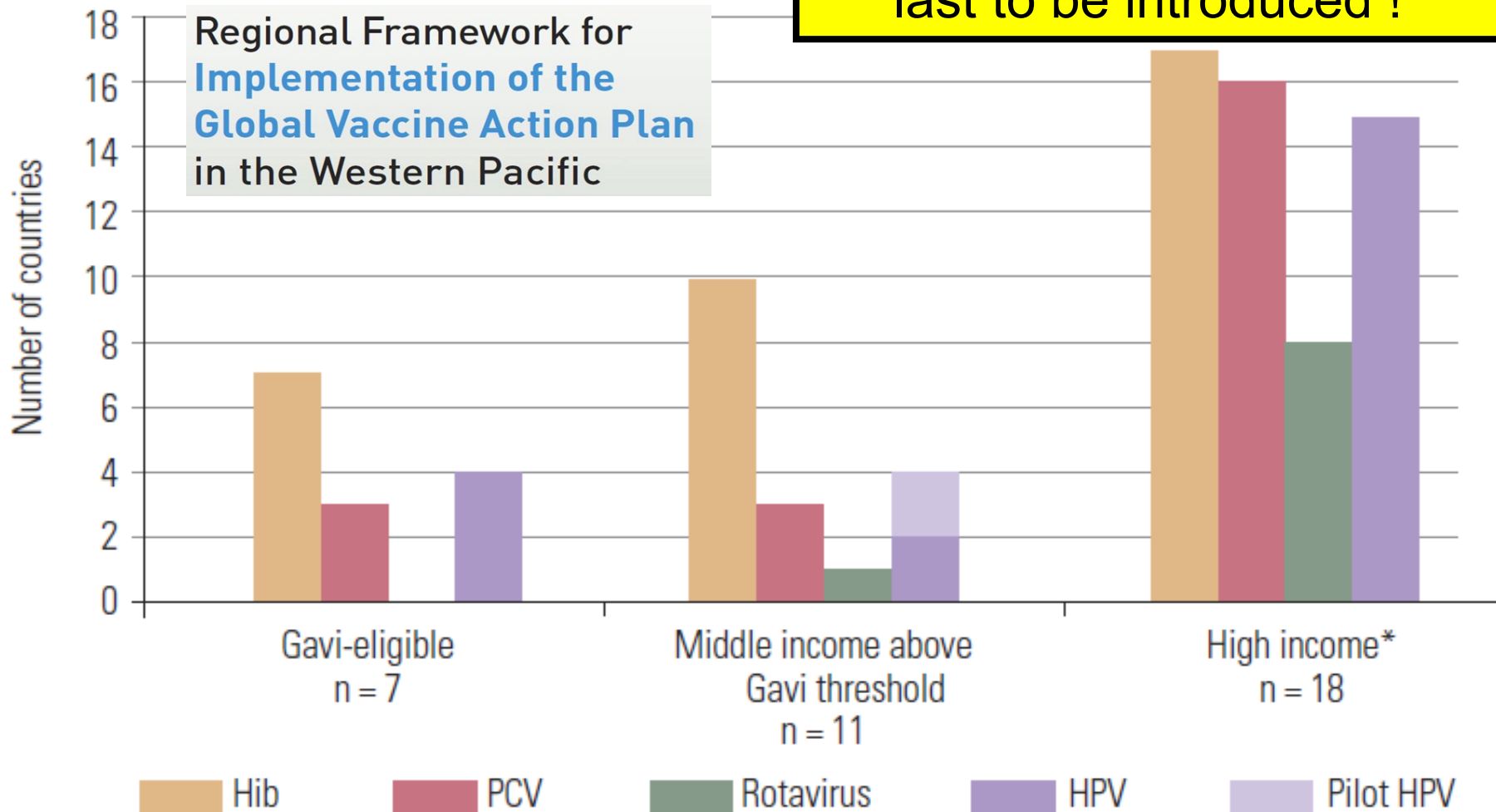


FIGURE 3. Countries and areas that have introduced new and underutilized vaccines, by income level, Western Pacific Region



*Including Pacific island countries and areas associated with the United States

Source: World Health Organization, Regional Office for the Western Pacific



Myint Htwe Minister of Health and Sport



Congratulations for your support for the EPI program and for putting Rotavirus and HPV on the schedule in 2020 !

Vaccine Hesitancy : The need to create awareness and demand

- RV is not serious
- RV can be easily treated
- RV is the pediatricians bread and butter
- Vaccine is not essential because there are few deaths here
- Vaccine is risky – for intussusception
- Natural immunity is better
- Vaccine cost is too high
- Too many other vaccines competing for funds

Recent Global Partnerships

The last case of natural Smallpox in the world, Smallpox
Ali Maow Maalin, Somalia, 1976



1980 Declaration of the
eradication of smallpox





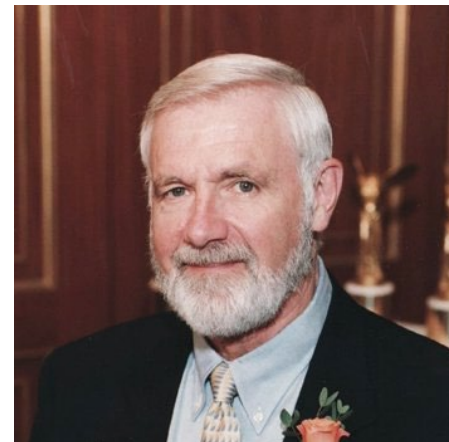
Meetings on Disease eradication Fogarty International Center, NIH 1980 & 1983



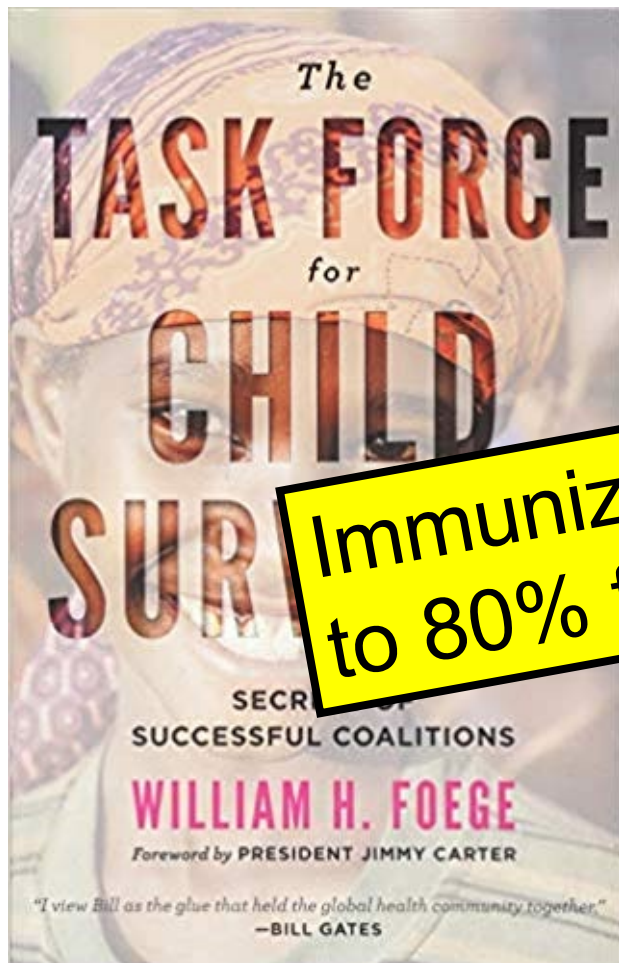
**END
POLIO
NOW**

**COUNTDOWN TO
HISTORY** 

Task force for child survival -1984-1990



Bill Foege
Lasker Awardee
2001



- 4000 preventable childhood deaths/day

Immunization rates increased to 80% for at least one vaccine!

How do we do better together?

UNICEF

WHO

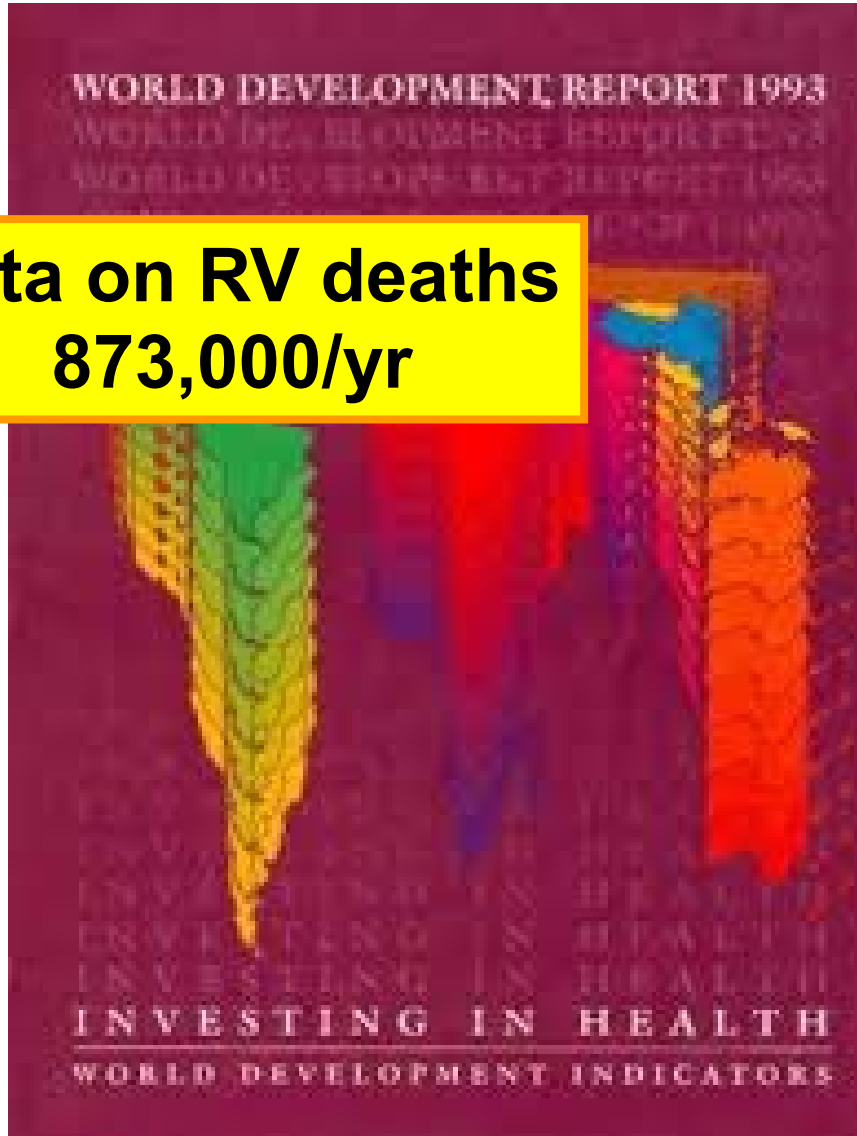
World Bank

UNDP

Rockefeller Foundation

World Bank Report 1993

**Data on RV deaths
873,000/yr**



Data on the
Global Burden of
disease

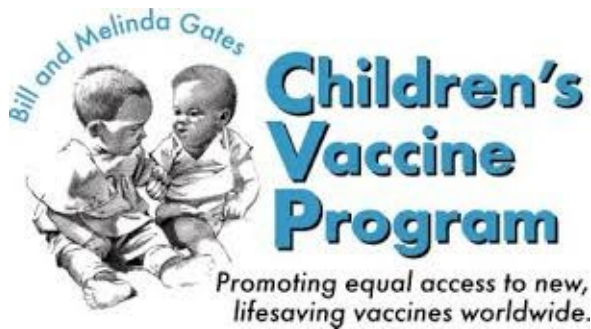
Why investing in
health makes
economic sense !

The Children's Vaccine Initiative

Achieving the Vision

- Goals -1990s
- Single dose
 - Heat stable
 - Multiple antigens
 - Research

Transitioned to GAVI
Global Alliance for
Vaccine Initiative



BILL & MELINDA
GATES *foundation*

1999

Accelerated Development & Introduction Plan

\$100 Million for Hep B, Hib Pneumo and RV Vaccines

- Accelerated Development and Introduction Plan (ADIP) established with generous grant from GAVI Alliance.
- Partnership with WHO and US CDC.
- Mission: *To reduce child mortality and morbidity from diarrheal disease by accelerating the availability of rotavirus vaccines appropriate for use in developing countries.*



How can we make vaccines affordable ?

“...[new] vaccines will be too expensive in the beginning to have developing countries buying them with their own governmental funds. But there is a renewed commitment from the international community to help.”

Dr Marie-Paule Kiery, Director, WHO Initiative for Vaccine Research, as quoted in UN health agency concerned over rising cost of vaccines for poor. *Agence France Presse*, 25 May 2005.

ADVANCE MARKET COMMITMENTS



New vaccines co-financing

Co-financing per dose (USD)

Vaccine	Examples	Co-financing per dose (USD)			Fragile States
		Poorest	Intermediate	Least Poor	
No. 1	1 st vaccine, single or combination vaccines (including yellow fever)	\$0.20	\$0.30	\$0.30	\$0.10
No. 2	2 nd additional vaccine (single or combination)	\$0.15	\$0.15	\$0.15 (+15% annually)	\$0.15
No. 3	3 rd additional vaccine (single or combination)	\$0.15	\$0.15	\$0.15 (+15% annually)	\$0.15

IFFIm

SUPPORTING GAVI

GAVI Vaccine Finance

The GAVI Alliance: Accelerating Africa's access to rotavirus vaccines



For more information:
gavialliance.org/support

Rotavirus kills more than half a million children a year

50% are in Africa

Angola
 Burundi
 Cameroun
 Congo
 Djibouti
 Ethiopia
 Ghana

Madagascar
 Malawi
 Niger
 Rwanda
 Sudan
 Tanzania

Benin
 Burkina Faso
 CAR
 Chad
 Comoros
 Congo DR
 Côte d'Ivoire
 Eritrea
 Gambia
 Guinea
 Guinea Bissau
 Kenya
 Lesotho

Liberia
 Mali
 Mauritania
 Mozambique
 Nigeria
 São Tomé
 Senegal
 Sierra Leone
 Togo
 Uganda
 Zambia
 Zimbabwe

Rotavirus vaccine support timeline

WHO recommends rotavirus vaccine be included in all national immunisation programs
 GAVI approve **\$4.5 Billion**

Replenish 2010

GAVI approves 12 more African countries for rotavirus vaccine support

\$7.0 Billion
Replenish 2015

25 more African countries are eligible for GAVI's rotavirus vaccine support

Country Price
\$ 0.15-.30/dose

vaccines to reach developing countries

The first GAVI-eligible country in Africa introduced the rotavirus vaccine two years

Rotavirus is the world's leading cause of

The New York Times

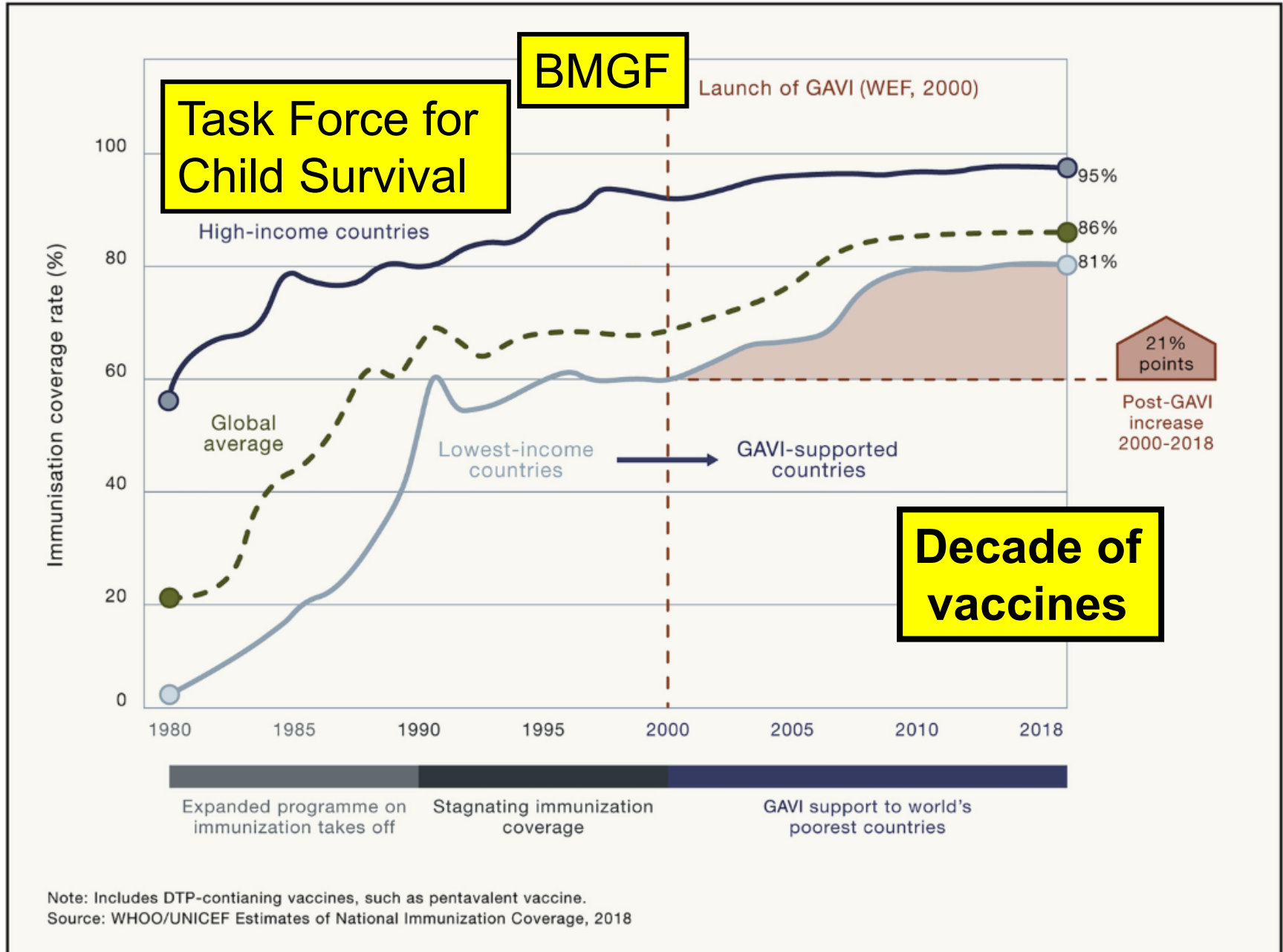
September 10, 2019

2019 Lasker-Bloomberg Public Service Award: GAVI, the Vaccine Alliance

for providing sustained access to childhood vaccines around the globe, saving millions of lives, and highlighting the power of immunization to prevent disease



Figure 1. Immunization Rates 1980–2018





DECADE *of*
VACCINES
COLLABORATION



The vision for the DoV is a world in which all individuals and communities enjoy lives free from vaccine-preventable diseases. Its mission is to extend, by 2020 and beyond, the full benefits of immunization to all people, regardless of where they are born, who they are, or where they live.





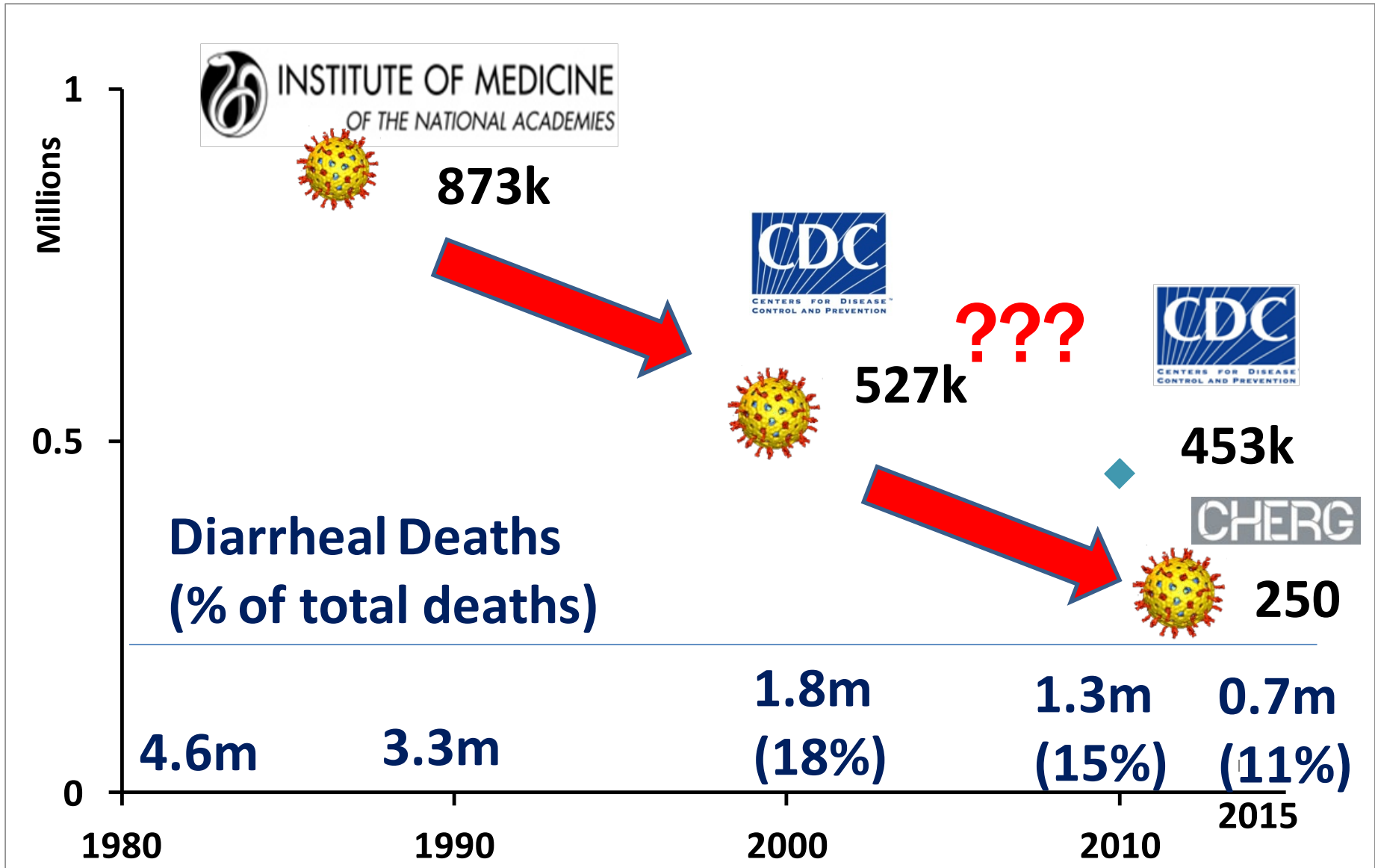
Where do we go next ?

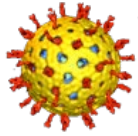
Gavi Board starts framing Alliance's approach to 2021-2025 period

Gavi Board approves in principle a set of new and expanded vaccine programmes.

- Hepatitis B birth dose (stop infection at birth & chronic hep B)
- DPT boosters at 12-24 mo, 4-7 years and 9-15 yrs.
- Oral cholera (OCV) –reduce incidence in poor, marginal groups
- Human rabies vaccine for post-exposure prophylaxis
- Meningococcal conjugate vaccine – A,C,W
- Respiratory Syncytial Virus – bronchiolitis/pneumonia infants

Estimates of Rotavirus Deaths: 2012





Rotavirus Vaccination and the Global Burden of Rotavirus Diarrhea Among Children Younger Than 5 Years

Christopher Troeger, MPH; Ibrahim A. Khalil, MD; Puja C. Rao, MPH; Shujin Cao, MS; Brigitte F. Blacker, MPH; Tahmeed Ahmed, MD; George Armah, PhD; Julie E. Bines, MD; Thomas G. Brewer, MD; Danny V. Colombara, PhD; Gagandeep Kang, MD; Beth D. Kirkpatrick, MD; Carl D. Kirkwood, PhD; Jason M. Mwenda, PhD; Umesh D. Parashar, MD; William A. Petri Jr, MD; Mark S. Riddle, MD; A. Duncan Steele, PhD; Robert L. Thompson, PhD; Judd L. Walson, MD; John W. Sanders, MD; Ali H. Mollad, PhD; Christopher J. L. Murray, DPhil; Simon I. Hay, FMedSci; Robert C. Reiner Jr, PhD



Chris Troeger

- 128 500 deaths (95% [UI], 104 500-155 600)
- 258 million episodes of diarrhea (95% UI, 193 million to 341 million),
- Incidence - 0.42 cases per child-year (95% UI, 0.30-0.53).
- Vaccine use averted > 28 000 deaths (95% UI, 14 600-46 700)
- Expanded vaccine use in sub-Saharan Africa, could have prevented approximately 20% of all deaths attributable to diarrhea among children

Final Thoughts --Partnerships & Collaborations

- The world recognizes that vaccines are among the most cost effective interventions in public health
- The vaccine community has come together –from the largest global agencies and donors to regional, national and local programs
- The number of new vaccines and variants is increasing challenging governments to consider each individually
- Anti-vaccine sentiment has grown and needs to be addressed
- Investments in research will be essential to address problems and make improvements over time



Mathuram Santosham
Johns Hopkins Bloomberg School
of Public Health, USA

Ciro A. de Quadros
Sabin Vaccine Institute, USA



Core Partners



Umesh Parashar
US Centers for Disease Control
and Prevention, USA

Kathy Neuzil
PATH, USA



Council Members

George Armah
University of Ghana, Ghana

Julie Bines
University of Melbourne, Australia



Shams El Arifeen
ICDDR, B, Bangladesh

Carlo Giaquinto
University of Padova, Italy



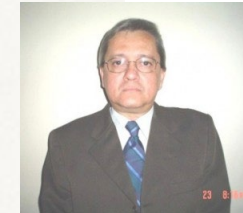
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Hospital, Malaysia



Gagandeep Kang
Christian Medical College, India

Alexandre C. Linhares
Ministry of Health, Brazil



Erkin Musabaev
Institute of Virology, Uzbekistan

Tony Nelson
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Kong, Hong Kong



Vesta Richardson
Ministry of Health, Mexico

Duncan Steele
Bill & Melinda Gates Foundation,



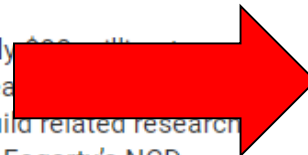
Oyewale Tomori
Redeemer's University, Nigeria

Anita Zaidi
Aga Khan University, Pakistan



Fogarty programs spur noncommunicable diseases research

Fogarty and its NIH partners invested nearly \$50 million to establish noncommunicable diseases research centers between U.S. and LMIC institutions, and build related research training programs according to a review of Fogarty's NCD programs between 2001 and 2017.



www.FIC.NIH.Gov

MORE >

Latest News

- ▶ March / April 2019 *Global Health Matters* newsletter

Funding News

- ▶ BLOODSAFE: Research to enhance availability of safe blood for patients with severe anemia and hemorrhagic conditions in low or lower-middle income countries (LLMICs) in Sub-Saharan Africa from NHLBI Due June 11 > Exploratory / Developmental Phased Award (RFA-HL-20-009) > Data Coordinating Center (RFA-HL-20-010)
- ▶ Global Infectious Disease Research Training Due July 25
- ▶ U.S.-South Africa Program for Collaborative Biomedical Research

Making a Difference



- ▶ Fogarty Fellow Dr Eric J Nelson's cholera research in Bangladesh with icddr, spawns global projects.



- ▶ Scientists inspired by a Fogarty workshop urge cross-cutting stigma research, interventions.

Fogarty Trainees

Fogarty's programs have made significant contributions by filling the pipeline of global health leaders. Featured researchers share how training supported by Fogarty has impacted their careers.

"Fogarty's aim is to develop researchers who can generate the evidence needed to affect policy, and be willing to step in and talk with policymakers about what their evidence shows."

Dr. Jonathan Samet, Dean, Colorado School of Public Health

- ▶ More featured Fogarty trainees

