



ACADEMIA
BRASILEIRA
DE CIÊNCIAS



UNIVERSIDADE
FEEVALE

Water and Human Health

Viral Gastroenteritis in Brazil

Gastroenterites Virais no Brasil

José Paulo Gagliardi Leite

Laboratory of Comparative and Environmental Virology



IOC
Oswaldo Cruz Institute



Brazilian Ministry of Health
FIOCRUZ
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Ministry of
Health

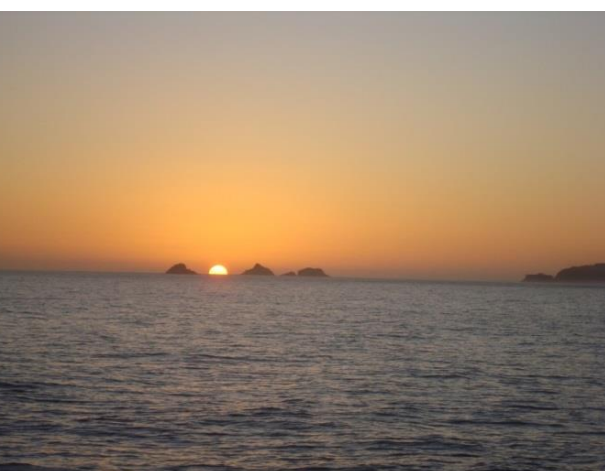
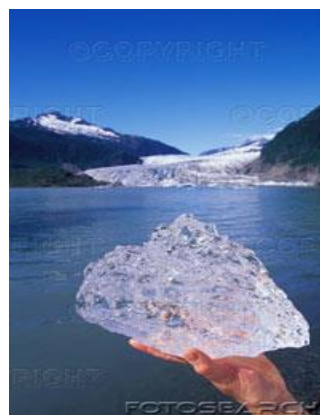
BRAZILIAN GOVERNMENT
BRASIL

Environmental, Cultural, Social and Health Sanitation Conditions





O QUE RECEBEMOS



O QUE DEVOLVEMOS



WATER

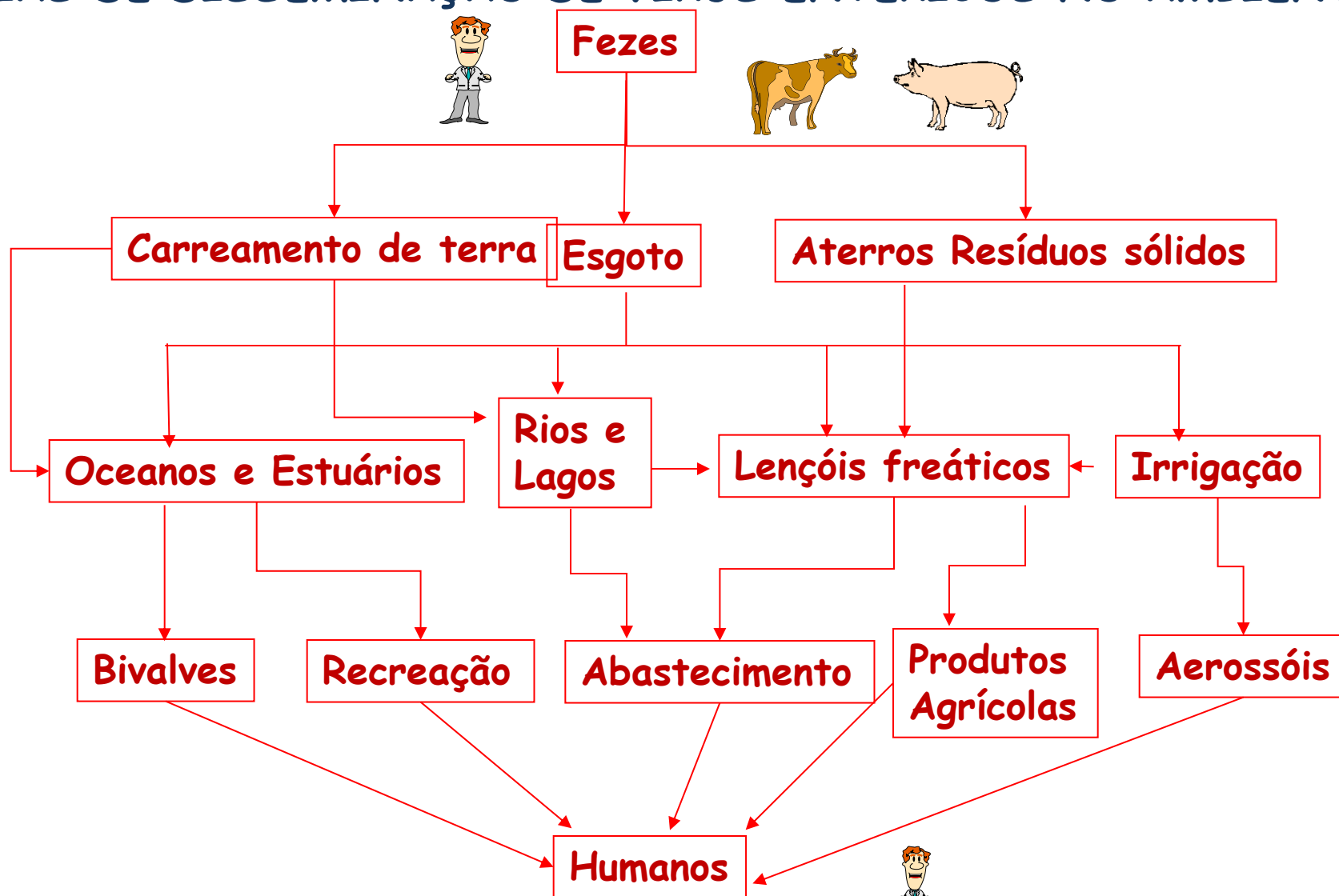
X

PUBLIC HEALTH



DE ONDE CHEGAM?

VIAS DE DISSEMINAÇÃO DE VÍRUS ENTÉRICOS NO AMBIENTE

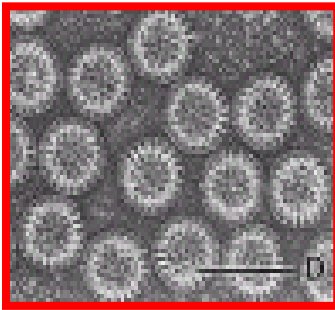


Effect of city-wide sanitation programme on reduction in rate of childhood diarrhoea in northeast Brazil: assessment by two cohort studies

Mauricio L Barreto^{a*}, Bernd Genser^a, Agostino Strina^a, Maria Gloria Teixeira, Ana Marlucia O Assis^c, Rita F Rego^d, Carlos A Teles^a, Matildes S Prado^a, Sheila MA Matos^a, Darci N Santos^a, Lenaldo A dos Santos^a, and Sandy Cairncross^b

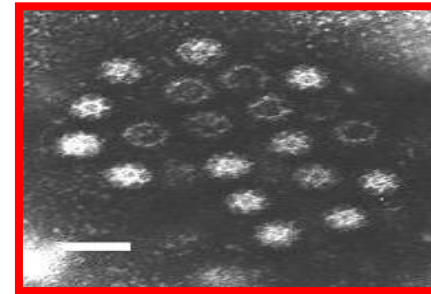
Findings—Diarrhoea prevalence fell by 21% (95% CI 18–25%)—from 9·2 (9·0–9·5) days per child-year before the intervention to 7·3 (7·0–7·5) days per child-year afterwards. After adjustment for baseline sewerage coverage and potential confounding variables, we estimated an overall prevalence reduction of 22% (19–26%).



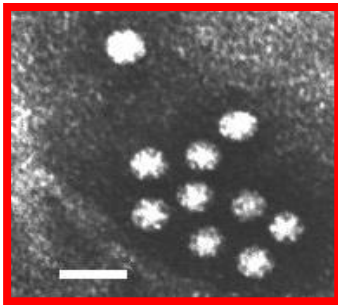


Rotaviruses

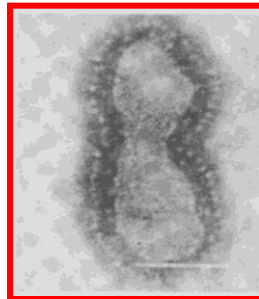
WATER, PUBLIC HEALTH and GASTROENTERIC VIRUSES



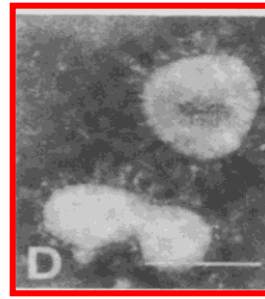
Human Caliciviruses



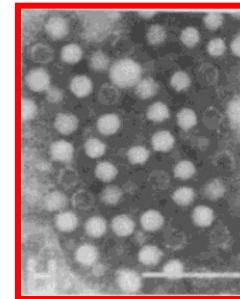
Astroviruses



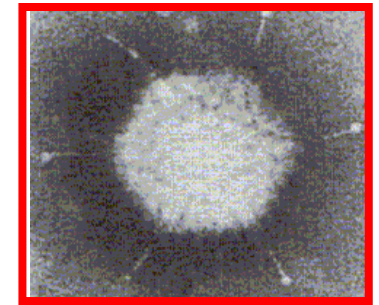
coronavirus



bredavirus



bocavirus



Enteric adenoviruses

Gut metagenomic analysis: new enteric viruses !!

12450–12455 | PNAS | July 23, 2013 | vol. 110 | no. 30

www.pnas.org/cgi/doi/10.1073/pnas.1300833110

Rapid evolution of the human gut virome

Samuel Minot^a, Alexandra Bryson^a, Christel Chehoud^a, Gary D. Wu^b, James D. Lewis^{b,c}, and Frederic D. Bushman^{a,1}

Fastidious!

Experimental model?

Interspecies infections !?

Genetic
diversity

Antigenic
diversity

Diversity
of
hosts





RISCO PARA SAÚDE PÚBLICA

1. GRANDE NÚMERO DE PARTÍCULAS VIRAIS ELIMINADAS NAS FEZES DE INDIVÍDUOS INFECTADOS (10^5 - $10^{11}/g$)

1 grama de fezes = 10.000.000 Vírus

1.000.000 Bactérias

1.000 Parasitas

100 Ovos de parasita

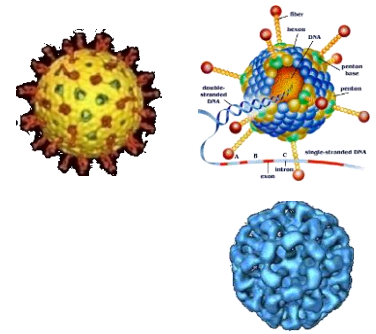
2. BAIXA DOSE INFECCIOSA - O CONTATO COM APENAS UMA ÚNICA PARTÍCULA VIRAL É CAPAZ DE DESENCADENAR UMA INFECÇÃO

10 a 10000 vezes > risco de bactérias

Haas & Gerba, 1991

3. ESTABILIDADE E RESISTENCIA

- NO ESGOTO: RESISTEM AOS TRATAMENTOS SECUNDARIOS
- NOS SEDIMENTOS: ADSORVEM-SE AOS SEDIMENTOS $\Rightarrow$ ALTA ESTABILIDADE
- NOS MOLUSCOS: ADSORVEM-SE AOS TECIDOS (BRÂNQUIAS E TRATO GASTROINTESTINAL) $\Rightarrow$ ALTA ESTABILIDADE



EMERGING INFECTIOUS DISEASES®

Enteric Viruses



August 2013



EMERGING INFECTIOUS DISEASES

August 2013



On the Cover

Catherine M. Howell (1892–1975)
Oyster Shuckers (1934)
Oil on canvas (101.4 cm x 91.6 cm)
Smithsonian American Art Museum,
Transfer from the US Department of
the Interior, National Park Service,
Washington, DC, USA

About the Cover p. 1354



Emergency Department Visit Data for Rapid Detection and Tracking of Norovirus Activity, United States..... 1214

B. Rha et al.

These data estimated the onset, peak, and
end of norovirus season within 4 weeks of
observed dates.

Aichi Virus in Sewage and Surface Water, the Netherlands..... 1222

W.J. Lodder et al.

High rates of detection indicate that this
virus is common in the human population in
this country.

Commentaries

Rapid Advances in Understanding Viral Gastroenteritis from Domestic Surveillance 1189

D.C. Payne and U.D. Parashar

Beyond Discovering the Viral Agents of Acute Gastroenteritis..... 1190

R.I. Glass

Perspectives

The New Global Health 1192

K.M. De Cock et al.

Strong public health institutions and networks
are needed to address evolving challenges.

Norovirus Disease in the United States..... 1198

A.J. Hall et al.

Findings support development of interventions
for this disease.

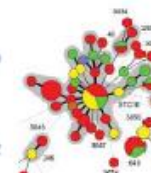
Research



Extended Spectrum β-Lactamase-producing *Enterobacteriaceae* among Travelers from the Netherlands 1206

S. Paltansing et al.

Colonization with potentially multidrug-resistant
bacteria was discovered at an elevated rate
among returning travelers.



p. 1210

p. 1236



Effects and Clinical Significance of GII.4 Sydney Norovirus, United States, 2012–2013..... 1231

E. Lesben et al.

Surveillance and emergency department data
showed that GII.4 Sydney did not cause a
substantial increase in norovirus activity.

Outbreak-associated *Salmonella* *enterica* Serotypes and Food Commodities, United States, 1998–2008..... 1239

B.R. Jackson et al.

Serotype–food commodity associations will aid
health officials in preventing illnesses.

Comparison of 2 Laboratory Assays for Diagnosis of Rotavirus in Children with Acute Gastroenteritis 1245

J.E. Tate et al.

The use of enzyme immunoassays is
recommended for evaluating vaccine
effectiveness.

Emerging Infectious Diseases • www.cdc.gov/eid • Vol. 19, No. 8, August 2013

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Brazilian Ministry of Health
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Oswaldo Cruz Foundation

Ministry of
Health





Rapid Advances in Understanding Viral Gastroenteritis from Domestic Surveillance

Daniel C. Payne and Umesh D. Parashar

COMMENTARY

Beyond Discovering the Viral Agents of Acute Gastroenteritis

Roger I. Glass

PERSPECTIVE

The New Global Health

Kevin M. De Cock, Patricia M. Simone, Veronica Davison, and Laurence Slutsker

PERSPECTIVE

Norovirus Disease in the United States

Aron J. Hall, Ben A. Lopman, Daniel C. Payne, Manish M. Patel, Paul A. Gastañaduy, Jan Vinjé, and Umesh D. Parashar

Travel-associated Diseases, Indian Ocean Islands, 1997–2010

Hélène Savini, Philippe Gautret, Jean Gaudart, Vanessa Field, Francesco Castelli, Rogelio López-Vélez, Poh Lian Lim, Marc Shaw, Frank von Sonnenburg, Louis Loutan, and Fabrice Simon,
for the GeoSentinel Surveillance Network¹

Data collected by the GeoSentinel Surveillance Network for 1,415 ill travelers returning from Indian Ocean islands during 1997–2010 were analyzed. Malaria (from Comoros and Madagascar), acute nonparasitic diarrhea, and parasitoses were the most frequently diagnosed infectious diseases. An increase in arboviral diseases reflected the 2005 outbreak of chikungunya fever.

“People are beginning to understand there is nothing in the world so remote that it can't impact you as a person.”

“As pessoas estão começando a entender que não existe nada no mundo, mesmo tão remoto, que não possa afeta-la como pessoa”

William H. Foege

***Director, US Centers for
Disease Control, 1977-1983***

Cláudio Paiva

claudio.paiva@oglobo.com.br

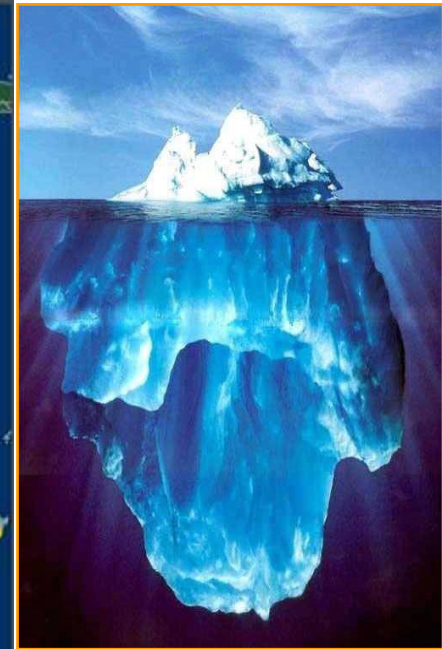
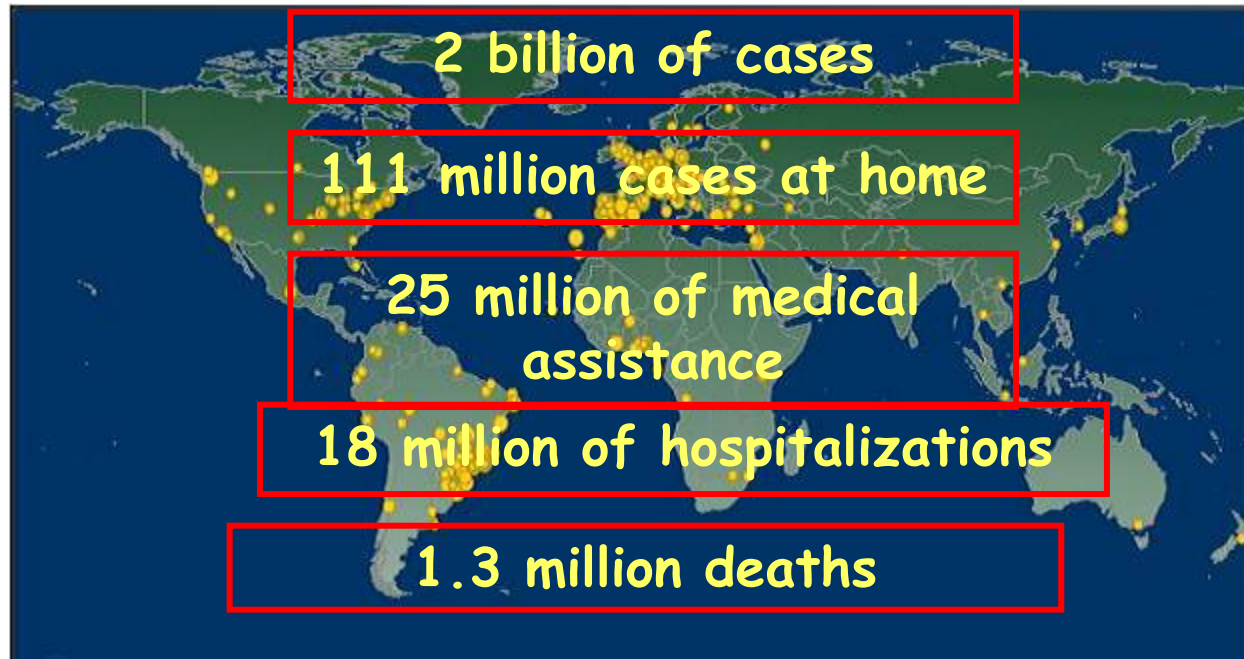
POR QUE GARÇONS NÃO DEVEM CUMPRIR AVISO PRÉVIO:



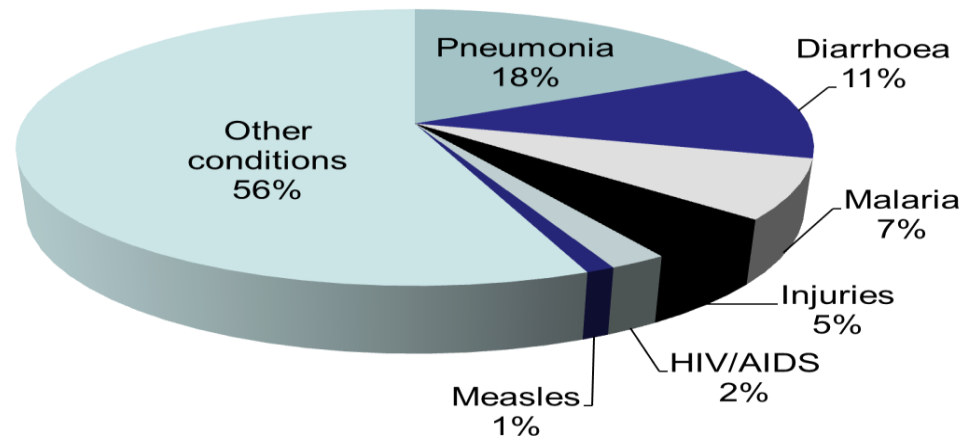
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www.CartoonStock.com

GASTROENTERITIS

MORBIDITY AND MORTALITY IN CHILDREN <5 YEARS OLD / YEAR



Parasites
Bacteria
viruses



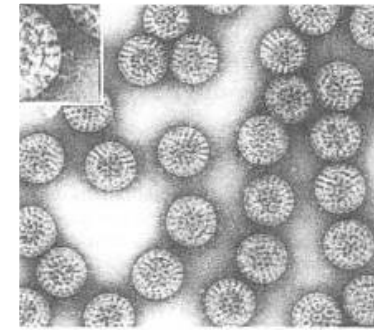
Aho *et al.* 2000; Mitchell, 2002; Kosek, 2003;
WHO 2005; Parashar *et al.*, 2007.

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BRAZILIAN GOVERNMENT
BRASIL



- **MOST COMMON CAUSE OF SEVERE DIARRHEA IN CHILDREN**
- **ALL CHILDREN WORLDWIDE INFECTED BY AGE 5**
- **STRAINS DIVERSITY CIRCULATION:**
DEVELOPING COUNTRIES ➤
DEVELOPED COUNTRIES
- **IMPROVEMENTS IN SANITATION WON'T SUBSTANTIALLY REDUCE DISEASE INCIDENCE : VACCINE !**

Major Clinical Symptoms
Diarrhea
Vomiting
Dehydration
Shock
Death

Burden and aetiology of diarrhoeal disease in infants and young children in developing countries (the Global Enteric Multicenter Study, GEMS): a prospective, case-control study

Karen L Kotloff, James P Nataro, William C Blackwelder, Dilruba Nasrin, Tamer H Farag, Sandra Panchalingam, Yukun Wu, Samba O Sow, Dipika Sur, Robert F Breiman, Abu S G Faruque, Anita K M Zaidi, Debasish Saha, Pedro L Alonso, Boubou Tamboura, Doh Sanogo, Uma Onwuchekwa, Byomkesh Manna, Thandavarayan Ramamurthy, Suman Kanungo, John B Ochieng, Richard Omore, Joseph O Oundo, Anowar Hossain, Sumon K Das, Shahnawaz Ahmed, Shahida Qureshi, Farheen Quadri, Richard A Adegbola, Martin Antonio, M Jahangir Hossain, Adebayo Akinsola, Inacio Mandomando, Tacilta Nhampossa, Sozinho Acácio, Kousick Biswas, Ciara E O'Reilly, Eric D Mintz, Lynette Y Berkeley, Khitam Muhsen, Halvor Sommerfelt, Roy M Robins-Browne, Myron M Levine

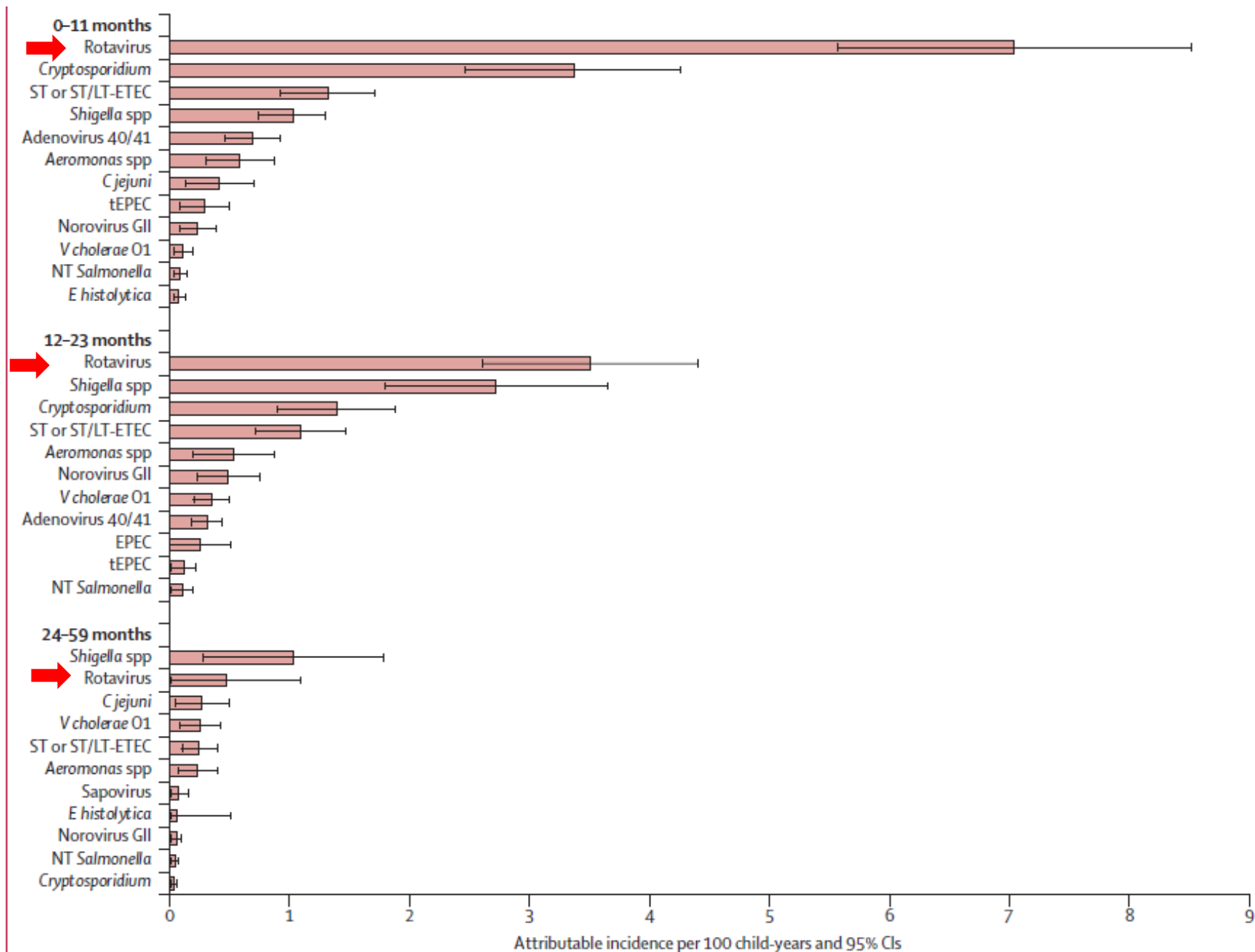
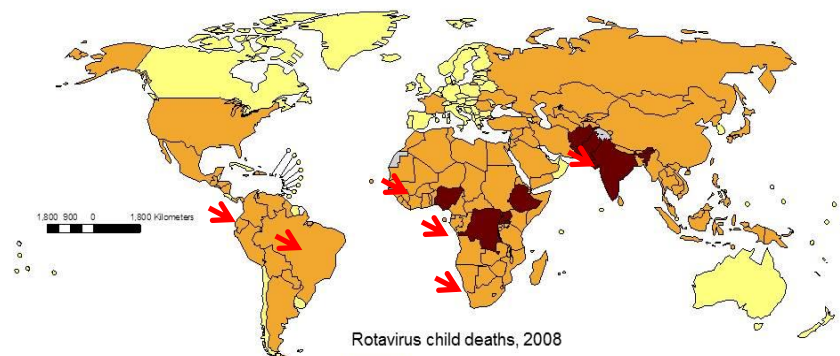
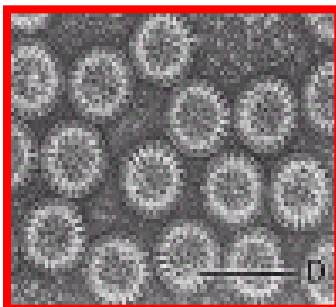


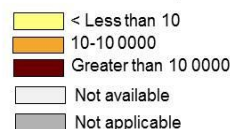
Figure 4: Attributable incidence of pathogen-specific moderate-to-severe diarrhoea per 100 child-years by age stratum, all sites combined

The bars show the incidence rates and the error bars show the 95% CIs.

453 000 global child rotavirus deaths, 2008



Rotavirus child deaths, 2008

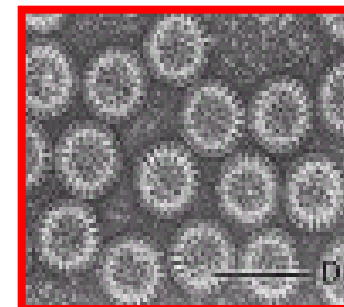


Data Source: WHO/IVB Rotavirus diseases burden estimates, January 2012

Map production: Immunization Vaccines and Biologicals, (IVB), World Health Organization

Date of slide: 02 February 2012

The boundaries and names shown and the designations used on this map do not imply the expression of any opinion whatsoever on the part of the World Health Organization concerning the legal status of any country, territory, city or sea or of its authorities, or concerning the delimitation of its frontiers or boundaries. Dotted lines on maps represent approximate border lines for which there may not yet be full agreement.
© WHO 2012. All rights reserved.



2 | TITLE from VIEW and SLIDE MASTER | February 21, 2012



1 death / 12 minutes

Table 3 Deaths due to rotavirus in children aged younger than 5 years, stratified by region

World bank Region	Rotavirus deaths	
	Total	Per 100,000 children younger than 5 y
Africa	236,519	155.8
Asia	144,095	39.6
Europe	167	0.4
Latin America and Caribbean	5,000	8.8
North America	3	0.0
Oceania	328	11.6

Aruna Chandran¹
Sean Fitzwater¹
Anjie Zhen²
Mathuram Santosham¹

Biologics: Targets & Therapy
29 July 2010

ROTAVIRUS CLASSIFICATION

Family: *Reoviridae*

Genus: *Rotavirus*

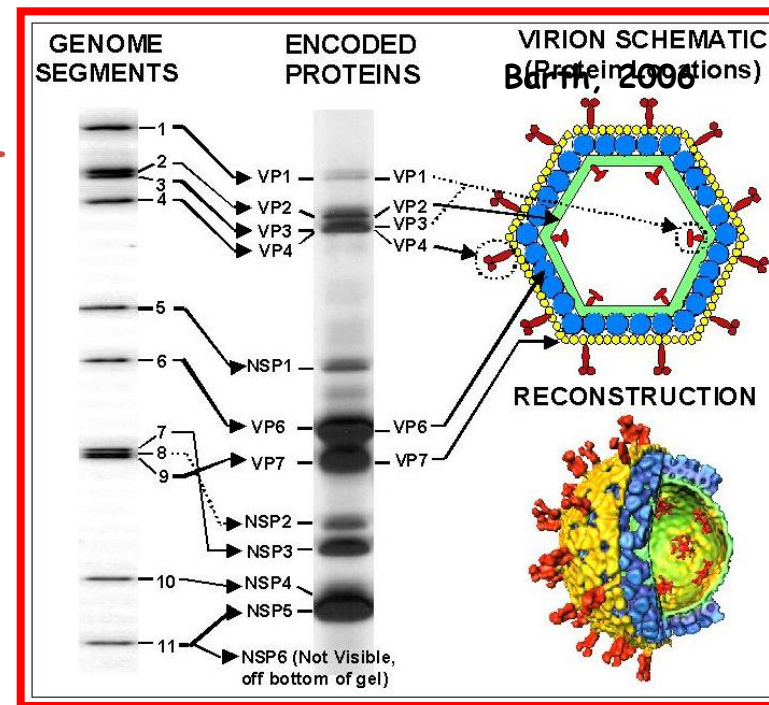
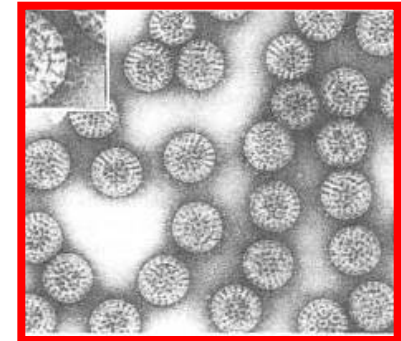
Species: **A** - G (H?) (Wakuda et al., 2011)

Virus Particle: 100 nm, non-enveloped, icosahedral with triple capsid protein.

Genome: dsRNA with 11 segments

Proteins: 6 structural (VP)
6 non-structural (NSP)

NSP4: enterotoxine
vomiting (Svensson 2009)



Modified from Estes, 2001 Estes & Kapikian, 2006

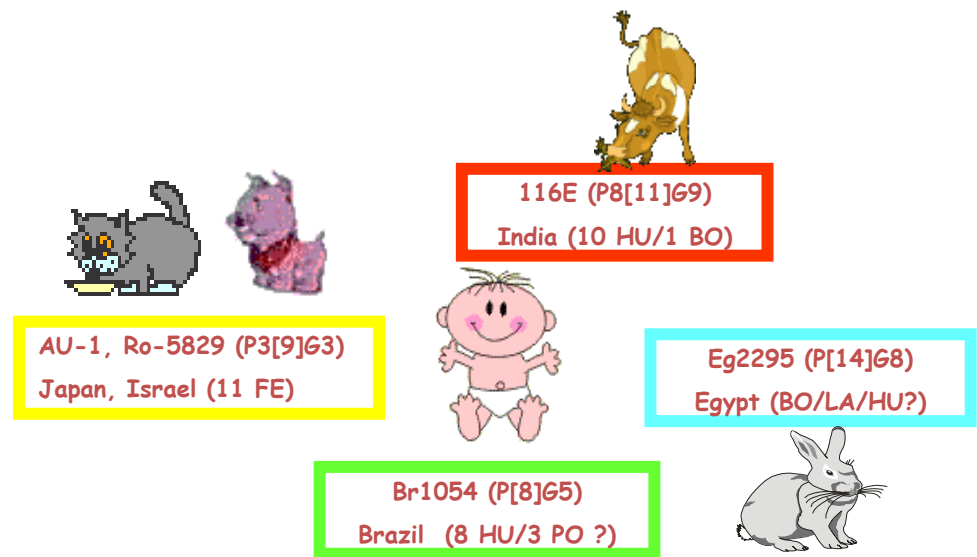
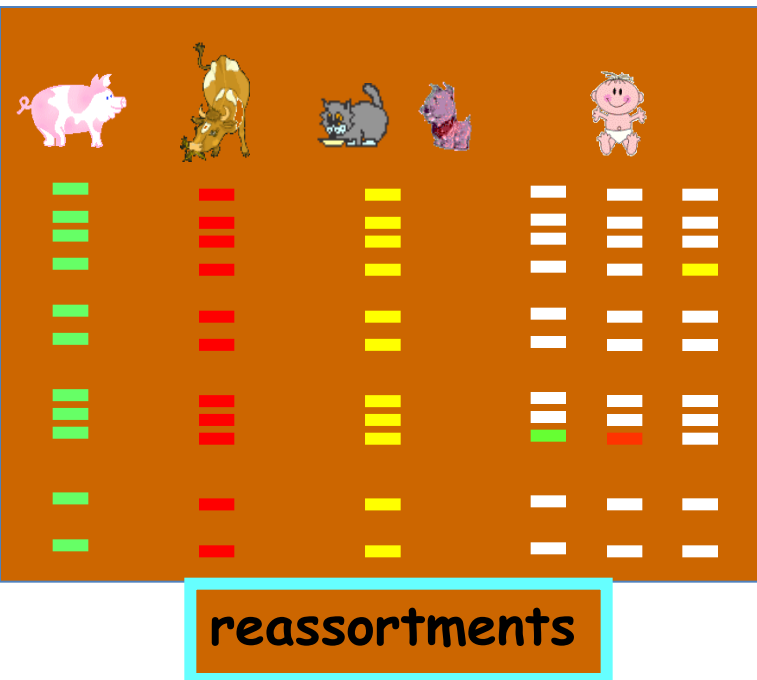
MECHANISMS FOR ROTAVIRUS EVOLUTION

Ponctual Mutation(s)

Rearrangements

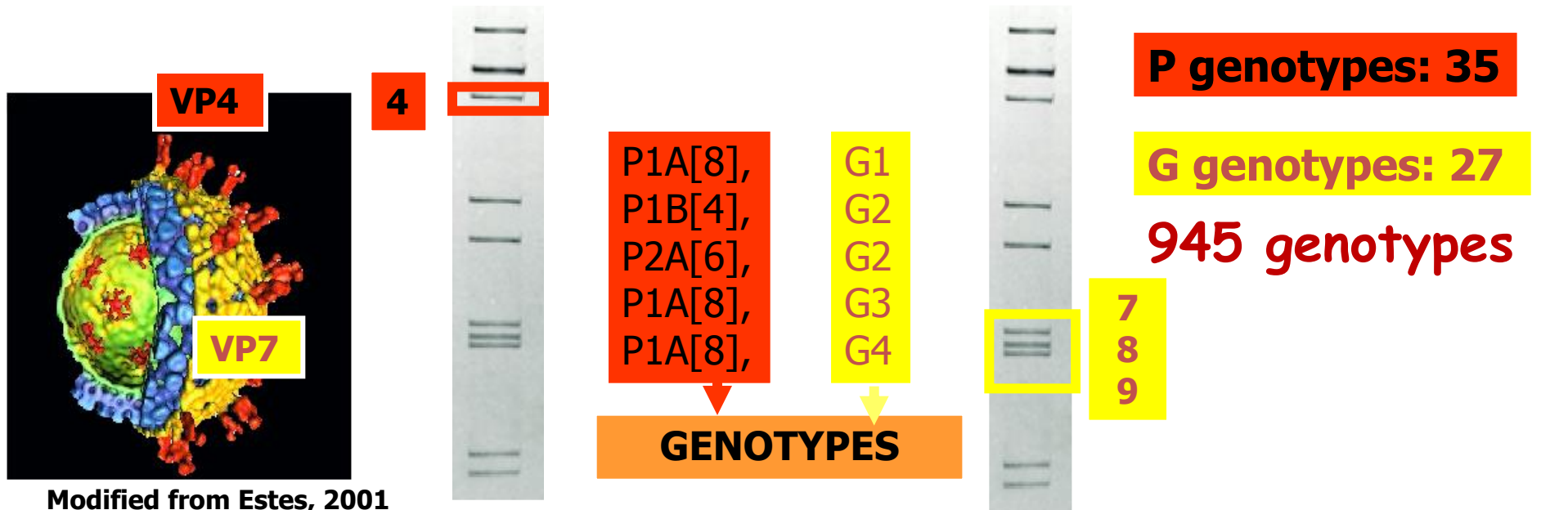
Reassortments

- Reassortment of RV-A strains
 - P and G antigen combinations
 - Intragenogroup reassortment of all genes
- Role of animal RV-A in diversity
 - Examples of interspecies infections
 - Human-animal RV-A reassortants



ROTAVIRUS SPECIE A

A BINARY CLASSIFICATION SYSTEM RVA: **P** and **G** GENOTYPES



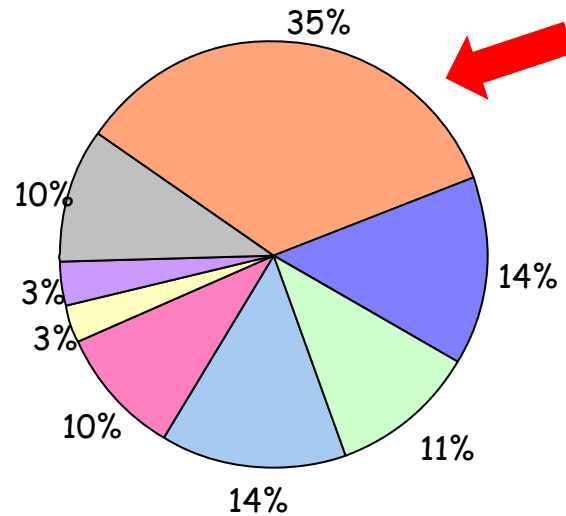
Prevalent worldwide human strains: **Wa-like:** P[8]G1, P[8]G3, P[8]G4, P[8]G9, and **DS-1 like:** P[4]G2

Prevalent in Asia and Africa human strains: G5, G8, G10, G12, P6, P9, P10

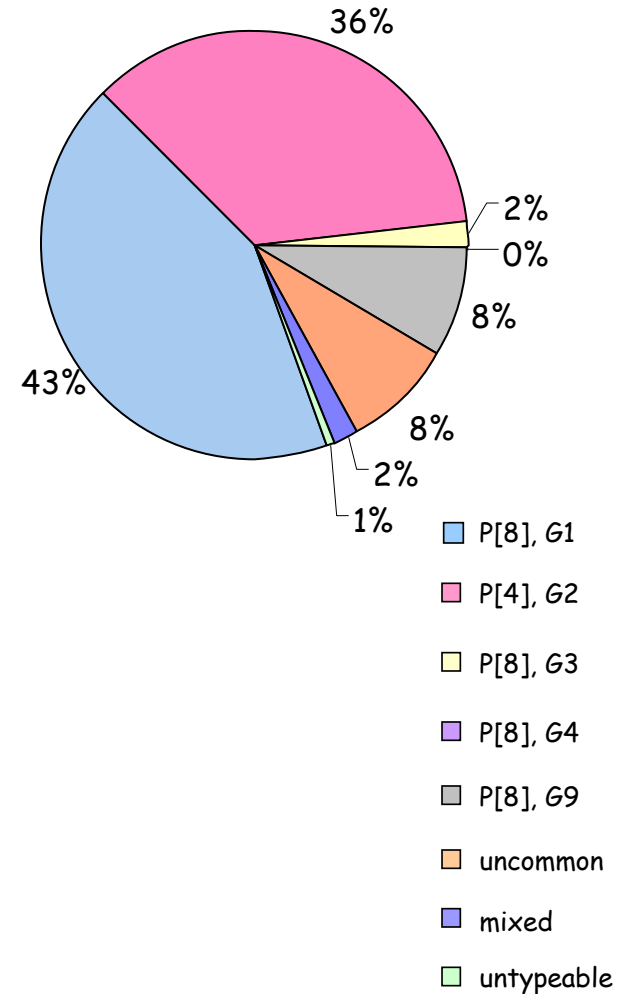
Khamrin et al., 2007; Martella et al., 2007; Matthijnssens et al., 2008, 2012; Albe et al., 2009

Distribution of Rotavirus Genotypes by WHO Regions, 2010

AFR, n=933

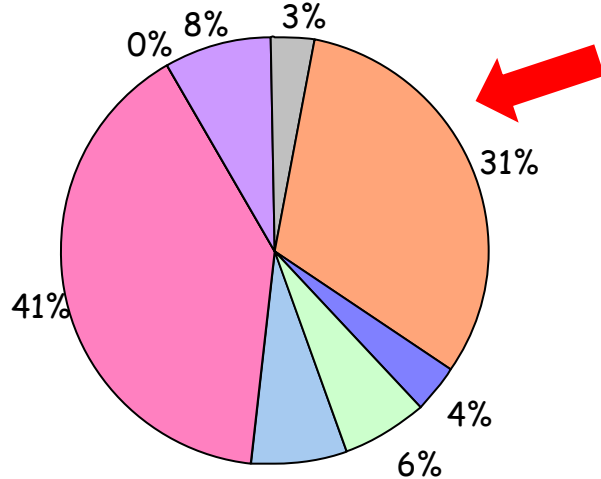


AMR, n=914



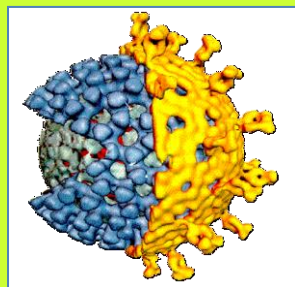
> Genotypes Diversity

EMR, n=702





Human Origin



G1P[8]

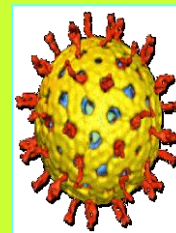
Rotarix® (GSK-Bio)



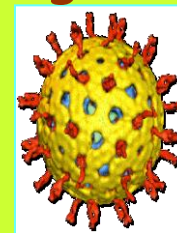
GSK - BioManguinhos / Fiocruz - Brazil



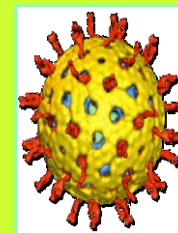
Human-Bovine Origin



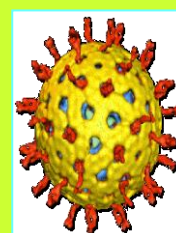
G1



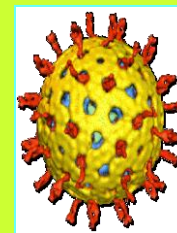
G3



P[8]



G2



G4



RotaTeq®
(Merck)



Review

Temporal trends in diarrhea-related hospitalizations and deaths in children under age 5 before and after the introduction of the rotavirus vaccine in four Latin American countries

L.H. De Oliveira^{a,d,*}, N. Giglio^{b,d}, A. Ciapponi^{c,d}, S. García Martí^{c,d}, M. Kuperman^{e,d}, N.J. Sanwogou^{a,d}, C. Ruiz-Matus^{a,d}, M.F. Marinho de Sousa^{a,d}

Table 2

Number of observed and expected deaths related to all-cause diarrhea following rotavirus vaccine introduction in children, by age group, in selected countries.

Country	Age range	Integral method			
		Observed deaths	Expected deaths	Difference	Δ% (CI 95%)
Bolivia	0–1 year	303	430 ± 4	127 ± 4	29.5% (28.2%–30.8%)
	0–5 years	314	487 ± 5	173 ± 5	35.5% (34.1%–36.9%)
El Salvador	0–1 year	188	178 ± 2	–10 ± 2	–5.6% (–8.0%––3.2%)
	0–5 years	253	240 ± 3	–13 ± 3	–5.4% (–8.3%––2.8%)
Honduras	0–1 year	85	99 ± 4	14 ± 4	14.1% (7.2%–21.1%)
	0–5 years	129	153 ± 2	24 ± 2	15.7% (13.5%–17.9%)
Venezuela	0–1 year	1010	2510 ± 10	1500 ± 10	59.8% (59.4%–60.1%)
	0–5 years	1535	3550 ± 32	2015 ± 32	56.8% (56.0%–57.5%)
Argentina	0–5 years	423	425 ± 1	–	0.4% (0.3%–0%)
Total ^a	0–1 year	1586	3217 ± 20	1631 ± 20	50.7% (50.1%–51.3%)
Total ^a	0–5 years	2231	4430 ± 42	2199 ± 42	49.6% (48.7%–50.6%)

DEATHS

Note: Deaths for the entire period, from the introduction of rotavirus vaccination through the last inf_{of}

^a The total does not include Argentina.

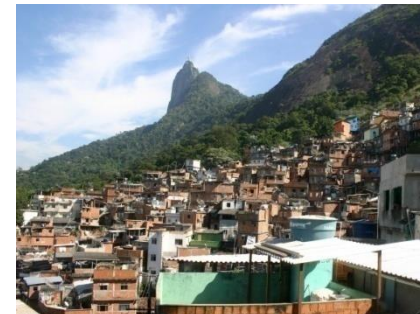
BRAZIL

- **Area: 8.514.876,599 km² (~40% LA)**
- **190.755.799 inhabitants**

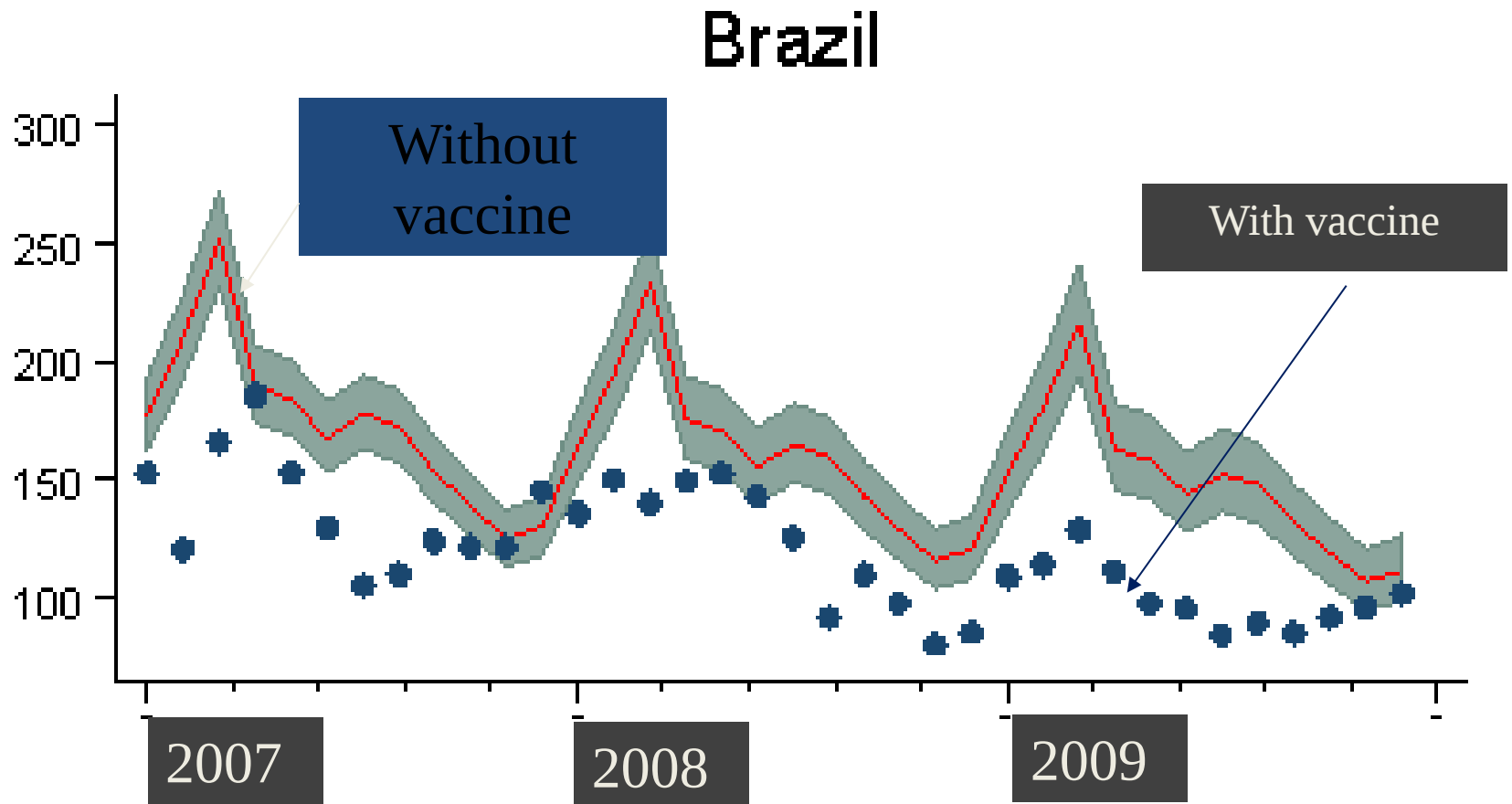


REGIONAL DIFFERENCES

Economical - Social
Environmental

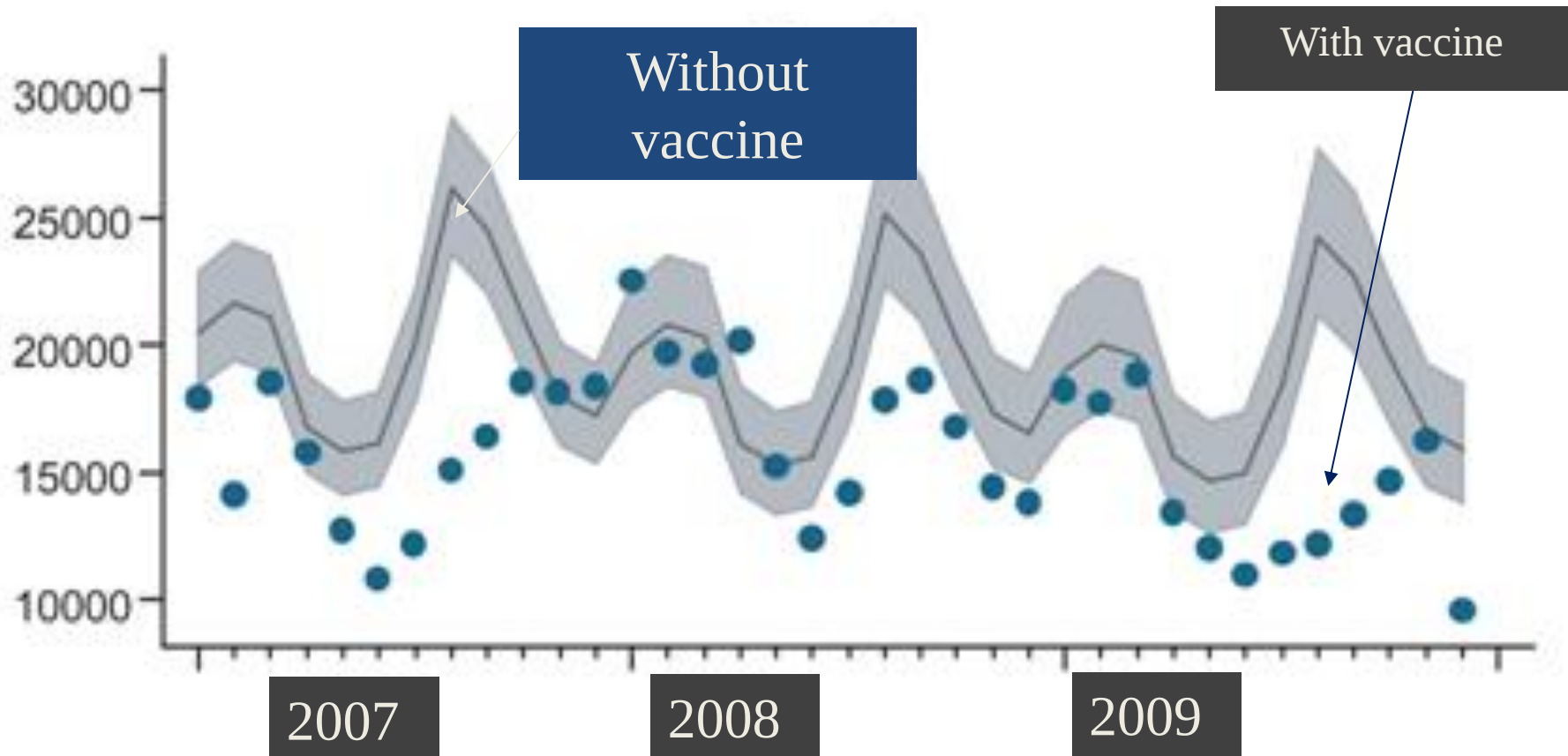


After the introduction of the Rotavirus vaccine , Brazil had 1500 fewer deaths under 5 due to diarrhea (2007-2009)



Carmo, G. et al. Plos Medicine, April 2011, Vol 8, Issue 4.

After Rotavirus vaccine introduction, Brazil had 130,000 fewer under 5 diarrhea admissions during 2007-2009

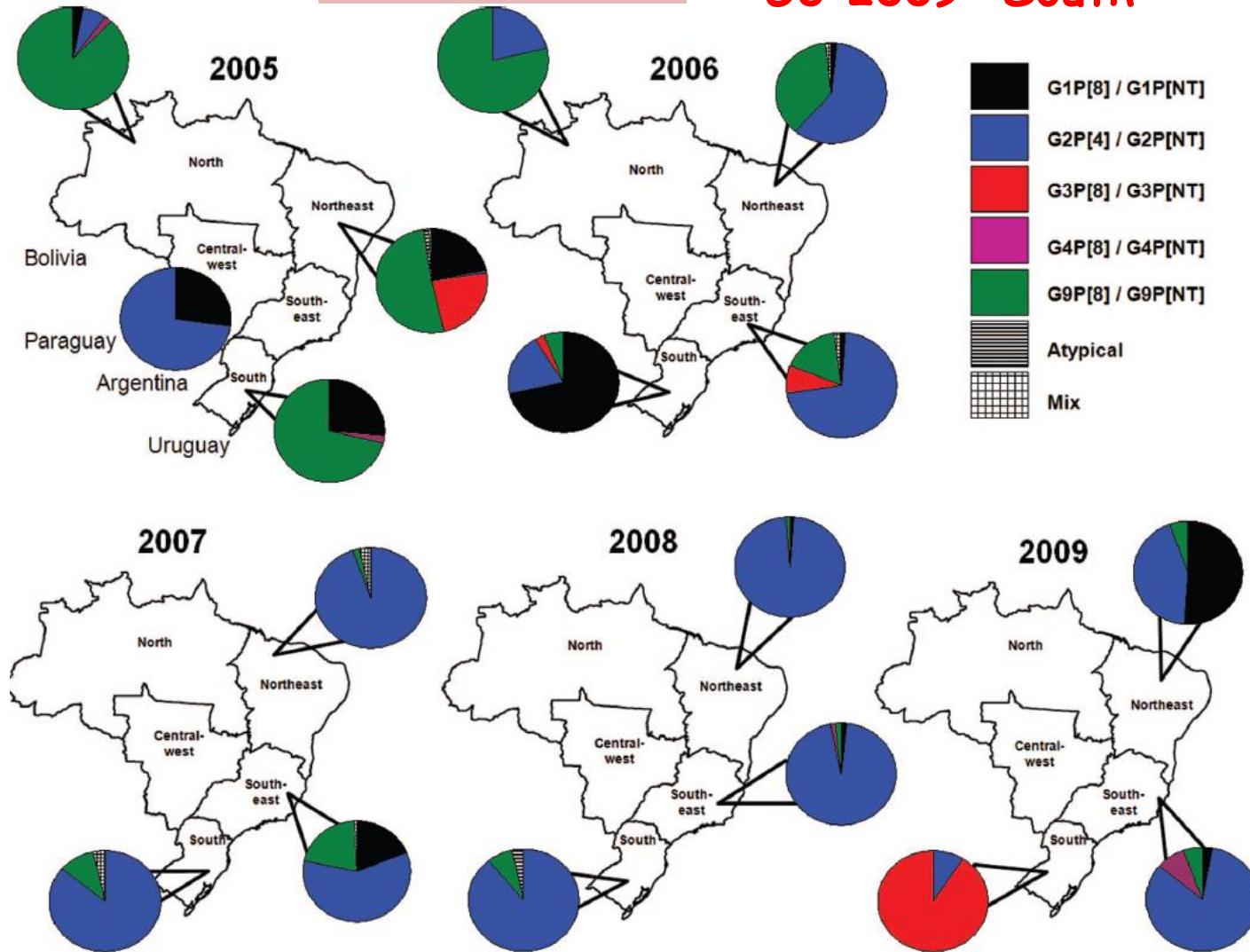


Carmo, G. et al. Plos Medicine, April 2011, Vol 8, Issue 4.

Filipe Anibal Carvalho-Costa, MD, PhD, Eduardo de Mello Volotão, PhD, Rosane Maria Santos de Assis, BSc, Alexandre Madi Fialho, BSc, Juliana da Silva Ribeiro de Andrade, BSc, Ludmila Nascimento Rocha, MSc, Luis Fernando López Tort, MSc, Marcelle Figueira Marques da Silva, MSc, Mariela Martinez Gómez, MSc, Pamella Macedo de Souza, BSc, and José Paulo Gagliardi Leite, PhD

Rotarix Introduction

G5 before 2005 / G9 after 2005
Prevalence of G2P[4] after 2005
G3 2009 South



Rotavirus surveillance, assessment of Rotarix (RV1) effectiveness and trends in diarrhea-associated mortality following massive vaccination in Brazil

Carvalho-Costa et al em preparação

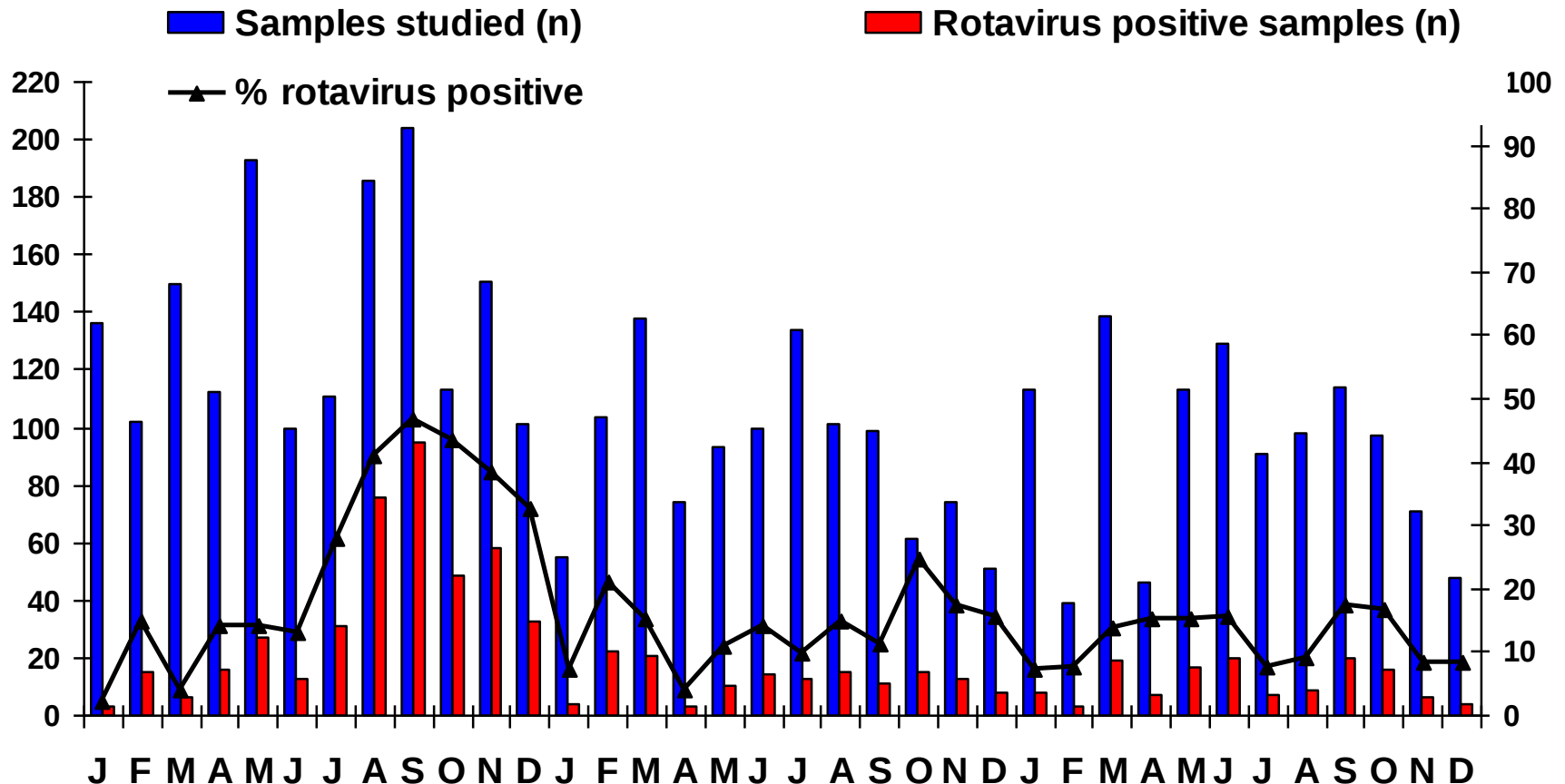


Figure 1. Monthly rates and seasonality of rotavirus detection in cases of acute diarrhea in Brazil, 2010 to 2012.

Conclusions:

RVA detection rate in subjects with acute diarrhea declined significantly from 2010 to 2012.

A clear shift in RVA genotypes distribution could be observed between 2010-2012.

This shift may have influenced positively RV1 performance due to reduction on prevalence of heterotypical strains.

RVA infection, presents a changing epidemiological profile in Brazil. RV1 vaccine, introduced in 2006, can be considered as one of several measures being implemented in Brazil with the goal of reducing diarrhea morbidity.

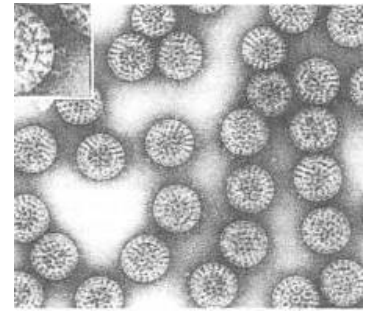
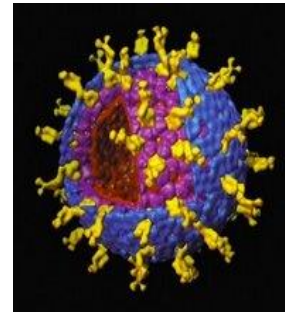
CHALLENGES RVA VACCINES



Biological Characteristics of RVA



Rotavirus Specie A



- **Infectivity: 10 -100 particles**
- **Shedding: 1 billion of infectious particles**
- **Environmental**

Emerging genotypes: G12, G8, G5, P6, P9 ? New vaccines?

JOURNAL OF CLINICAL MICROBIOLOGY, June 2006, p. 2046–2050 Vol. 44, No. 6

Molecular Epidemiology of Group A Rotavirus Diarrhea among Children in Buenos Aires, Argentina, from 1999 to 2003 and Emergence of the Infrequent Genotype G12

A. A. Castello,^{1,2*} M. H. Argüelles,² R. P. Rota,² A. Olthoff,³ B. Jiang,¹ R. I. Glass,¹ J. R. Gentsch,¹ and G. Glikmann²

Rotavirus infections with multiple emerging genotypes in Sri Lanka

Kamruddin Ahmed · Ranjith Batuwanthudawe · T. G. A. Nilmini Chandrasena · Marcelo Takahiro Mitui · Shaman Rajindrajith · Geethani Galagoda · Sher Bahadur Pun · Ryuichi Uchida · Osamu Kunii · Kazuhiko Moji · Nihal Abeysinghe · Akira Nishizono · Osamu Nakagomi

Genetic characterization of rotavirus subtypes in Pakistan-first report of G12 genotype from Pakistan under WHO-Eastern Mediterranean region

Muhammad Masroor Alam^a, Salman Akbar Malik^b, Shahzad Shaukat^a, Asif Naeem^a, Salmaan Sharif^a, Mehar Angez^a, Muhammad Suleman Rana^a, Adnan Khurshid^a, Sohail Zahoor Zaidi^{a,*}

Characterization of Genotype P[9]G12 Rotavirus Strains From Argentina: High Similarity With Japanese and Korean G12 Strains

Alejandro A. Castello,^{1,2*} Toyoko Nakagomi,³ Osamu Nakagomi,³ Baoming Jiang,² Jung O. Kang,⁴ Roger I. Glass,² Graciela Glikmann,¹ and Jon R. Gentsch²

National Rotavirus Surveillance in Argentina: High incidence of G9P[8] strains and detection of G4P[6] strains with porcine characteristics

Juan A. Stupka^a, Paola Carvalho^a, Alberto A. Amarilla^b, Mario Massana^c, Gabriel I. Parra^{b,*} the Argentinean National Surveillance Network for Diarrheas¹

Secular Variation in United States Rotavirus Disease Rates and Serotypes

Implications for Assessing the Rotavirus Vaccination Program

Daniel C. Payne, PhD, MSPH,* Peter G. Szilagyi, MD, MPH,† Mary Allen Staat, MD, MPH,‡ Kathryn M. Edwards, MD,§ Jon R. Gentsch, PhD,¶ Geoffrey A. Weinberg, MD,† Caroline B. Hall, MD,† Aaron T. Curns, MPH,* Haley Clayton, MPH,*|| Marie R. Griffin, MD, MPH,** Gerry Fairbrother, PhD,‡ and Umesh D. Parashar, MB BS, MPH*

Arch Virol (2012) 157:135–139
DOI 10.1007/s00705-011-1111-z

BRIEF REPORT

Identification of lineage III of G12 rotavirus strains in diarrheic children in the Northern Region of Brazil between 2008 and 2010

Luana da Silva Soares · Patrícia dos Santos Lobo · Joana D'Arc Pereira Mascarenhas · Darivaldo Luz Neri · Sílvia de Fátima dos Santos Guerra · Alessilva do Socorro Lima de Oliveira · Régis Piloni Maestri · Darleise de Souza Oliveira · Euzeni Maria de Fátima Costa de Menezes · Alexandre da Costa Linhares

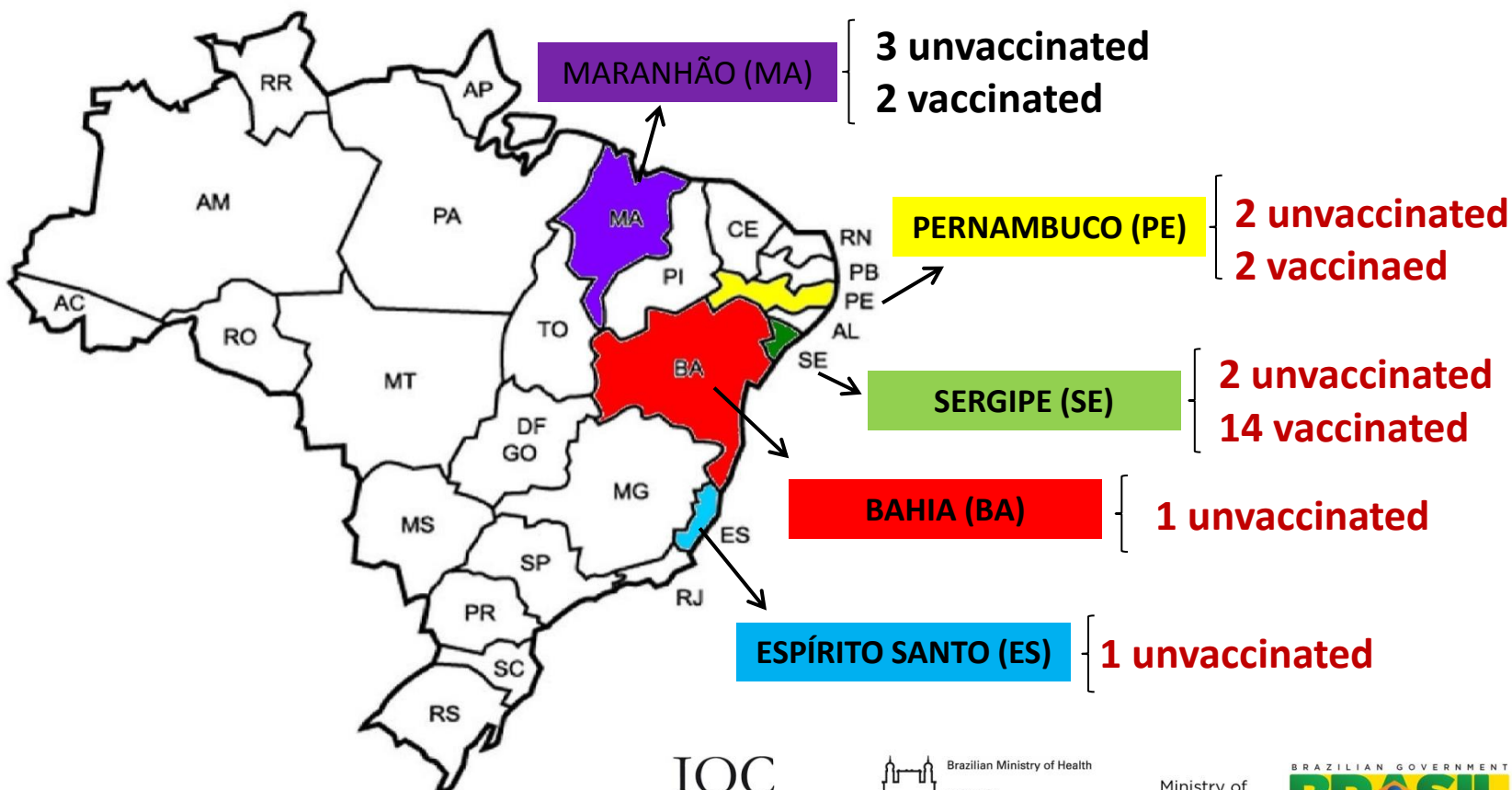
5 sentinel surveillance hospitals / 1 for each region

- 27 G1 P[8] < 5 years old
- 18 vaccinated / 9 unvaccinated
- May 2008 –August 2010

18 vaccinated from Northeastern

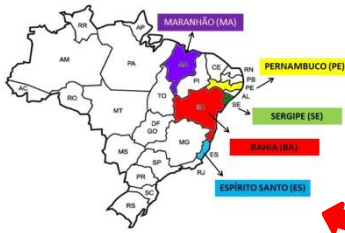
08 unvaccinated from Northeastern

1 unvaccinated from Southeastern



Intra-genotypic reassortment

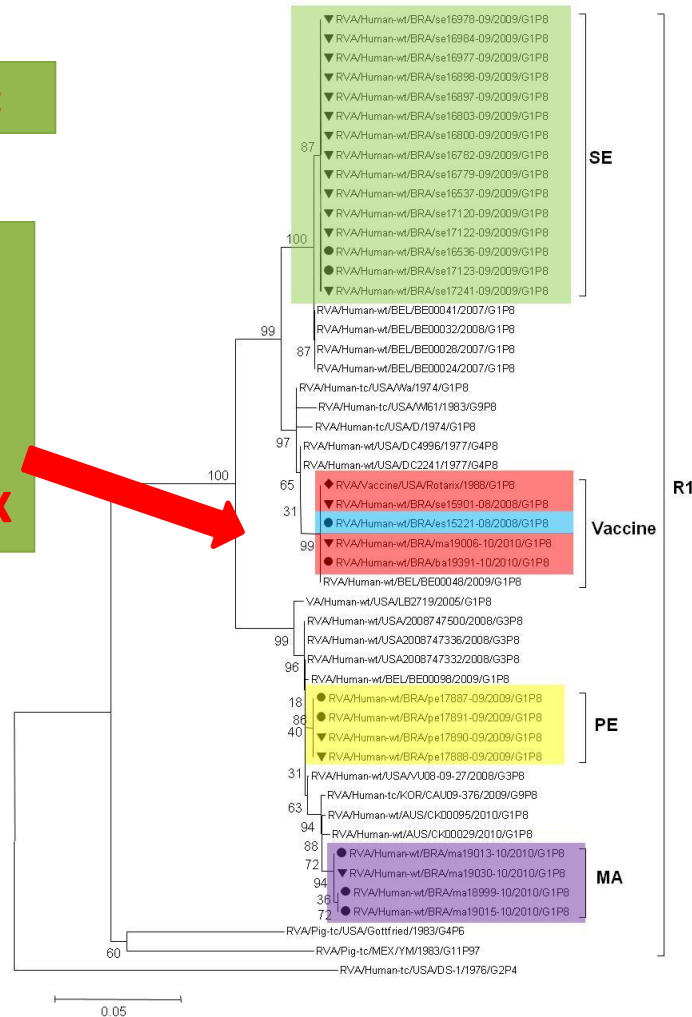
(VP1)



Southeast

VP2

**unvaccinated
child
42 months
VP1 from Rotarix**





Phylogenetic analysis of G1P[6] group A rotavirus strains detected in Northeast Brazilian children fully vaccinated with Rotarix™

Mariela Martínez Gómez ^{a,*}, Marcelle Figueira Marques da Silva ^a, Mark Zeller ^c, Elisabeth Heylen ^c, Jelle Matthijnsens ^c, Maria Yuri Travassos Ichihara ^b, Tatiana Lundgren Rose ^a, Eduardo de Mello Volotão ^a, Jose Paulo Gagliardi Leite ^a

Table 2

Genotype constellation comparison. The six Brazilian G1P[6] RVA strains analyzed in the current study are represented by RVA/Human-wt/BRA/BA17290/2009/G1P[6] strain. Green and red indicate Wa-like and DS-1-like gene segments, respectively. The P[6] VP4 genotype is colored yellow, and blue is used to indicate a gene segment of porcine origin. Brazilian strains are shown in bold.

	Strain	Origin	Genotypes										
			VP7	VP4	VP6	VP1	VP2	VP3	NSP1	NSP2	NSP3	NSP4	NSP5
Wa-like	RVA/Human-tc/USA/Wa/1974/G1P[8]	Human	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/USA/Rotarix-A41CB052A/1988/G1P[8]	Human	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-wt/BGD/Dhaka12/2003/G12P[6]	Human	G12	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-wt/ZAF/3176WC/2009/G12P[6]	Human	G12	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-wt/BGD/Matlab13/2003/G12P[6]	Human	G12	P[6]	I1	R1	C1	M1	A1	N1	T2	E1	H1
	RVA/Human-tc/KOR/CAU195/200X/G12P[6]	Human	G12	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-wt/DEU/GER172/2008/G12P[6]	Human	G12	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-wt/ECU/EC2184/200x/G11P[6]	Human	G11	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-tc/GBR/ST3/1975/G4P2A[6]	Human	G4	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-tc/AUS/RV3/1977/G3P[6]	Human	G3	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
	RVA/Human-wt/BRA/BA17290/2009/G1P[6]	Human	G1	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1
DS-1-like	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	Human	G2	P[4]	I2	R2	C2	M2	A2	N2	T2	E2	H2
	RVA/Human-wt/BEL/F01322/2009/G3P[6]	Human	G3	P[6]	I2	R2	C2	M2	A2	N2	T2	E2	H2
	RVA/Human-wt/COD/DRC86/2003/G8P[6]	Human	G8	P[6]	I2	R2	C2	M2	A2	N2	T2	E2	H2
	RVA/Human-wt/BGD/N26/2002/G12P[6]	Human	G12	P[6]	I2	R2	C2	M2	A2	N1	T2	E6	H2
	RVA/Human-wt/BGD/RV161/2000/G12P[6]	Human	G12	P[6]	I2	R2	C2	M2	A2	N2	T2	E1	H2
	RVA/Human-wt/BGD/RV176/2000/G12P[6]	Human	G12	P[6]	I2	R2	C2	M2	A2	N2	T2	E6	H2
	RVA/Human-wt/USA/06-242/2006/G2P[6]	Human	G2	P[6]	I2	R2	C2	M2	A2	N2	T2	E2	H2

relationship with contemporary typical human Wa-like RVA strains. These results suggests that the fact that these strains were able to cause acute gastroenteritis in vaccinated children is likely not due to the genetic background of the strains, but rather to other factors such as host-related factors or infection pathogens or vaccine efficacy. P[6] RVA strains



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Table 1 – Rotavirus specie A (RVA) detection from influent (24) and effluent (24) sewage samples by quantitative (qPCR) and qualitative (NSP4, VP4, VP7) PCR protocols for genotyping.

Year	Month	Sewage influent				Sewage effluent			
		qPCR	NSP4	VP4	VP7	qPCR	NSP4	VP4	VP7
2009	Aug	+	–	–	G2	–	–	–	–
		+	+	P[4]	G2	+	–	–	–
	Sep	+	+	P[4]	G2	+	–	–	–
		+	+	–	G2	+	–	–	–
	Oct	+	+	P[6]	G2	+	–	–	G2
2010		+	+	P[4]	G2	+	+	P[4]	G2
	Nov	+	+	P[6]	G2	+	+	–	G2
		+	+	P[4]	G2	+	–	–	–
	Dec	+	+	P[4]	G2	+	–	–	–
		+	+	P[6]	G2	–	–	–	–
2010	Jan	+	+	P[6]	G2	+	–	–	–
		+	+	–	G2	+	–	–	–
	Feb	+	+	P[6]	G2	+	–	–	–
		+	+	–	G2	–	–	–	–
	Mar	+	+	–	G2	–	–	–	–
		+	+	–	G2	+	+	–	G2
	Apr	+	+	–	G2	+	+	–	G2
		+	+	–	G2	+	–	–	G2
	May	+	+	P[4]	G2	–	–	–	–
		+	+	–	G2	+	–	–	–
	Jun	+	–	–	G2	+	–	–	–
		+	+	P[4]	G2	–	–	–	–
	Jul	+	+	P[6]	G2	–	–	–	–
		+	+	P[4]	G2	+	–	–	–

+: Positive; and –: negative.

One year environmental surveillance of rotavirus specie A (RVA) genotypes in circulation after the introduction of the Rotarix[®] vaccine in Rio de Janeiro, Brazil

Tulio Machado Fumian^{a,*}, José Paulo Gagliardi Leite^a, Tatiana Lundgreen Rose^a, Tatiana Prado^b, Marize Pereira Miagostovich^a

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BRASIL

Molecular Detection and Characterization of Gastroenteritis Viruses Occurring Naturally in the Stream Waters of Manaus, Central Amazônia, Brazil^V

Marize P. Miagostovich,¹ Fabiana F. M. Ferreira,¹ Flávia R. Guimarães,¹ Túlio M. Fumian,¹
 Leonardo Diniz-Mendes,¹ Sérgio Luiz B. Luz,² Luciete A. Silva,²
 and José Paulo G. Leite^{1*}

TABLE 2. Frequencies of the viruses investigated and fecal coliforms detected in the areas studied

Pathogen	No. (%) found in:		Chi square	P value
	Urban area (n = 24)	Rural and forest areas (n = 28)		
Any enteric virus	21 (87.5)	10 (35.7)	14.4	<0.001
RV	15 (62.5)	8 (28.6)	6.0	0.014
NoV	3 (12.5)		3.7	0.09
HAsTV	6 (25.0)	2 (7.1)	3.2	0.08
HAdV	15 (62.5)	1 (3.6)	21.1	<0.001
Fecal coliforms	24 (100.0)	14 (50.0)	16.4	<0.001

RV = HAdV > HAsTV > NoV

Detection of rotavirus A in sewage samples using multiplex qPCR and an evaluation of the ultracentrifugation and adsorption-elution methods for virus concentration

Tulio M. Fumian^{a,*}, José Paulo G. Leite^a, Alejandro A. Castello^b, Aldo Gaggero^c, Maria Susana L. de Caillou^d, Marize P. Miagostovich^a

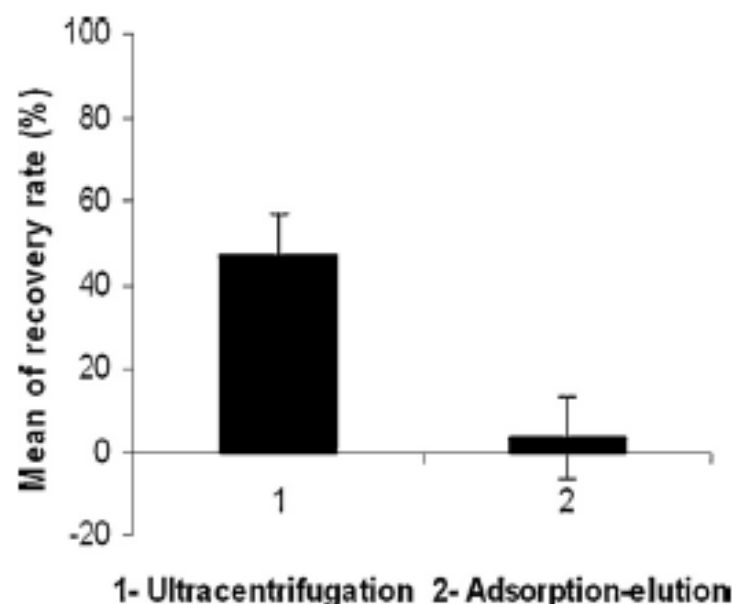
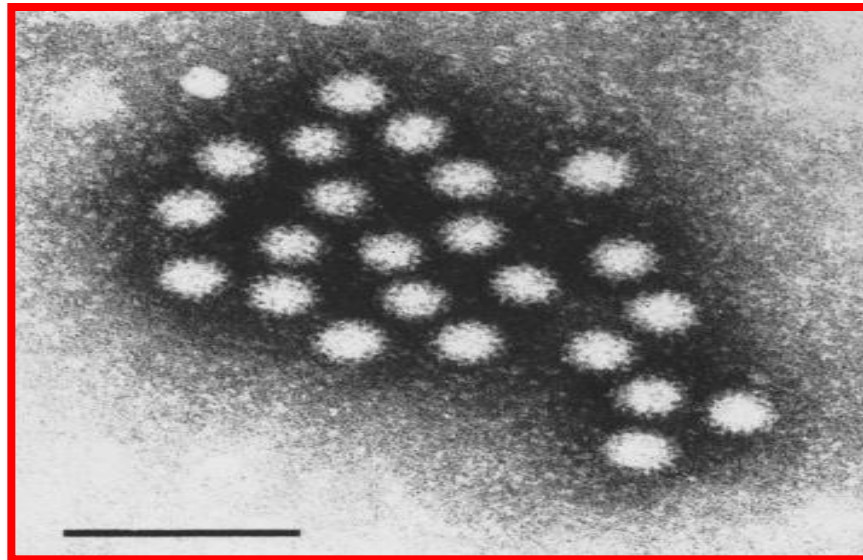


Fig. 1. The mean recovery rate of RV-A from sewage water following concentration using either the ultracentrifugation or adsorption-elution method ($n = 6$).

Burden of Norovirus infections

Patel et al. *Emerg. Infect. Dis.* 2008, 14(8):1224-31.

- **Developed countries:** ~900,000 episodes of diarrhea.
~64,000 hospitalization of children <5 years.
- **Developing countries:** ~1.1 million of hospitalizations
~218,000 deaths/year in children.



Electronic immunemicroscopy.

Kapikian. *J Infect Dis.* 2000 ,181(2): S295-302..

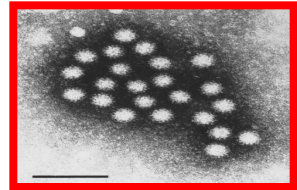
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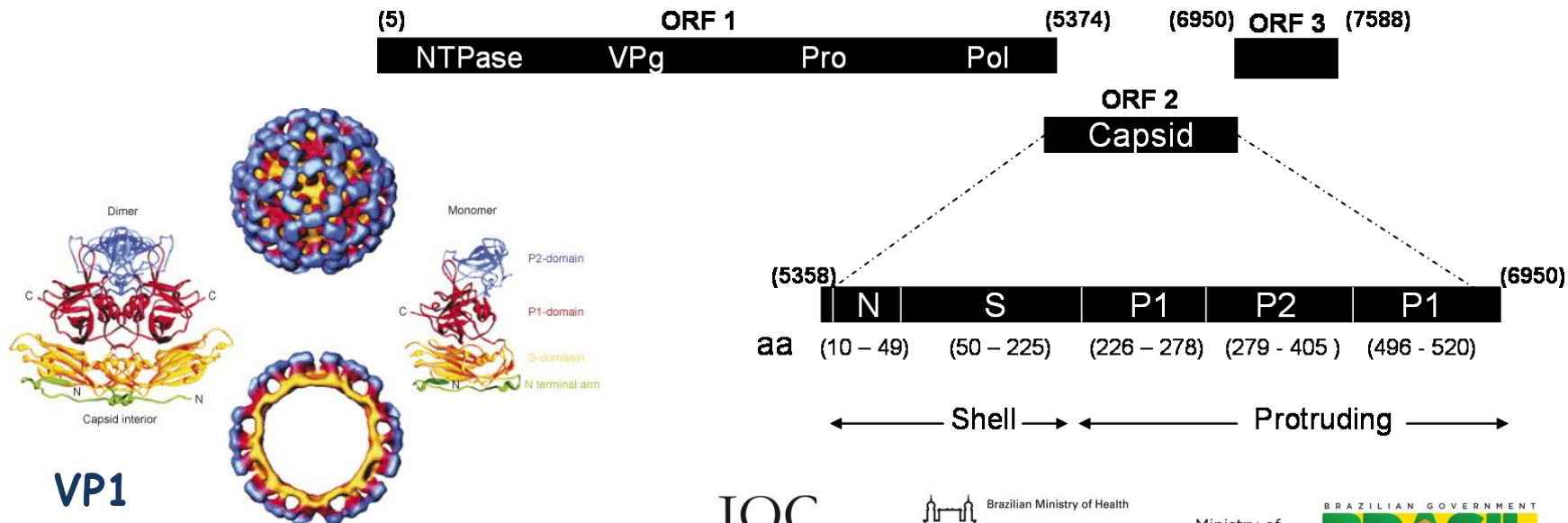
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Norovirus properties



- Non-enveloped virus, 38-40 nm in diameter
- Icosahedral symmetry, 180 VP1
- Single stranded positive RNA
- Non-cultivable
- Virus receptor seems to be an HBGA
- Replication in cytoplasm
- Exit probably by Cell lysis

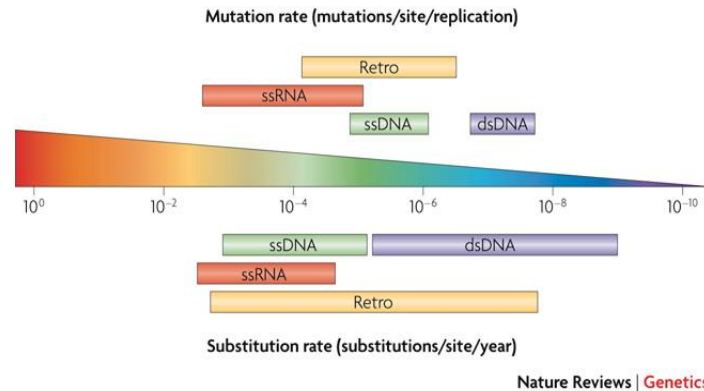


MECHANISMS FOR NOROVIRUS EVOLUTION

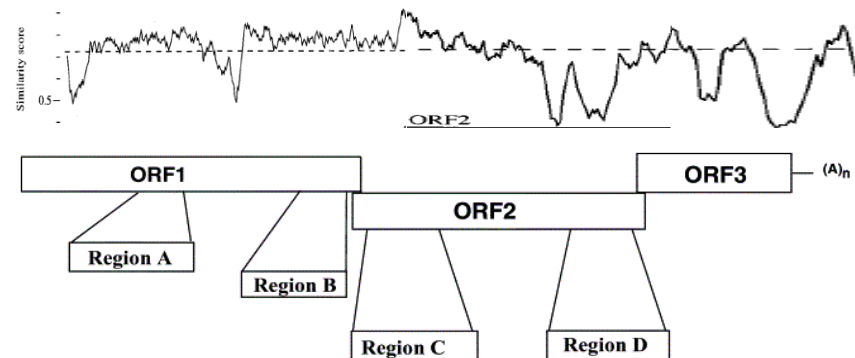
- **Ponctual Mutation(s)**

estimated mutation rate:

1.21×10^{-2} to 1.41×10^{-2} substitutions per site per year



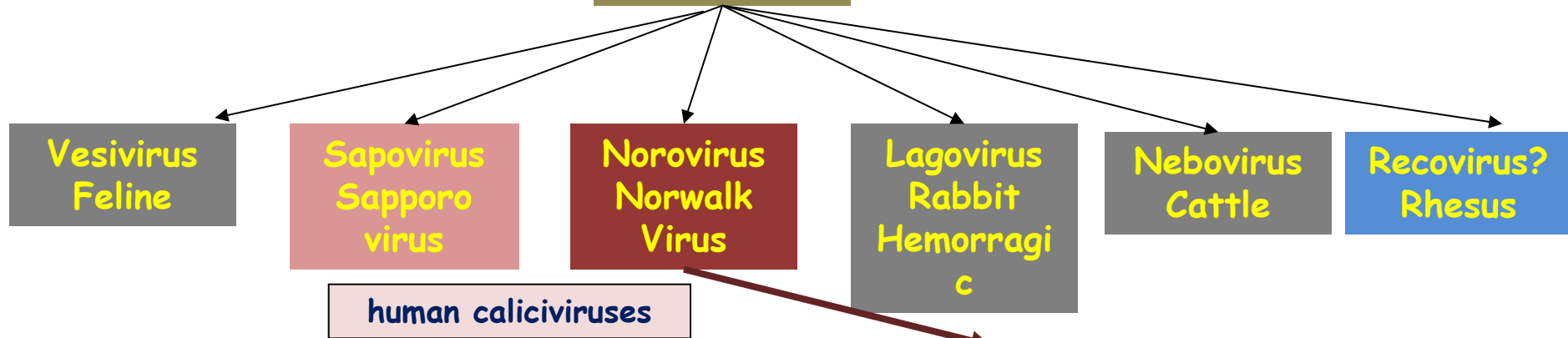
- **Recombination**



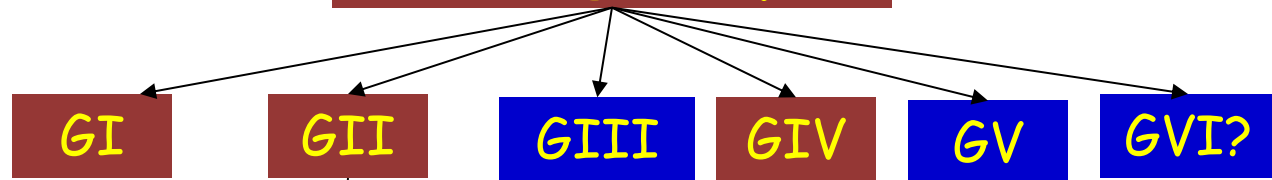
Victoria et al., Bayesian coalescent inference reveals high evolutionary rates and expansion of Norovirus populations. *Infect Genet Evol.* 2009 ; [Rates of evolutionary change in viruses: patterns and determinants](#). Duffy S, Shackelton LA, Holmes EC. *Nat Rev Genet.* 2008 Apr;9(4):267-76.

Caliciviridae

Genus



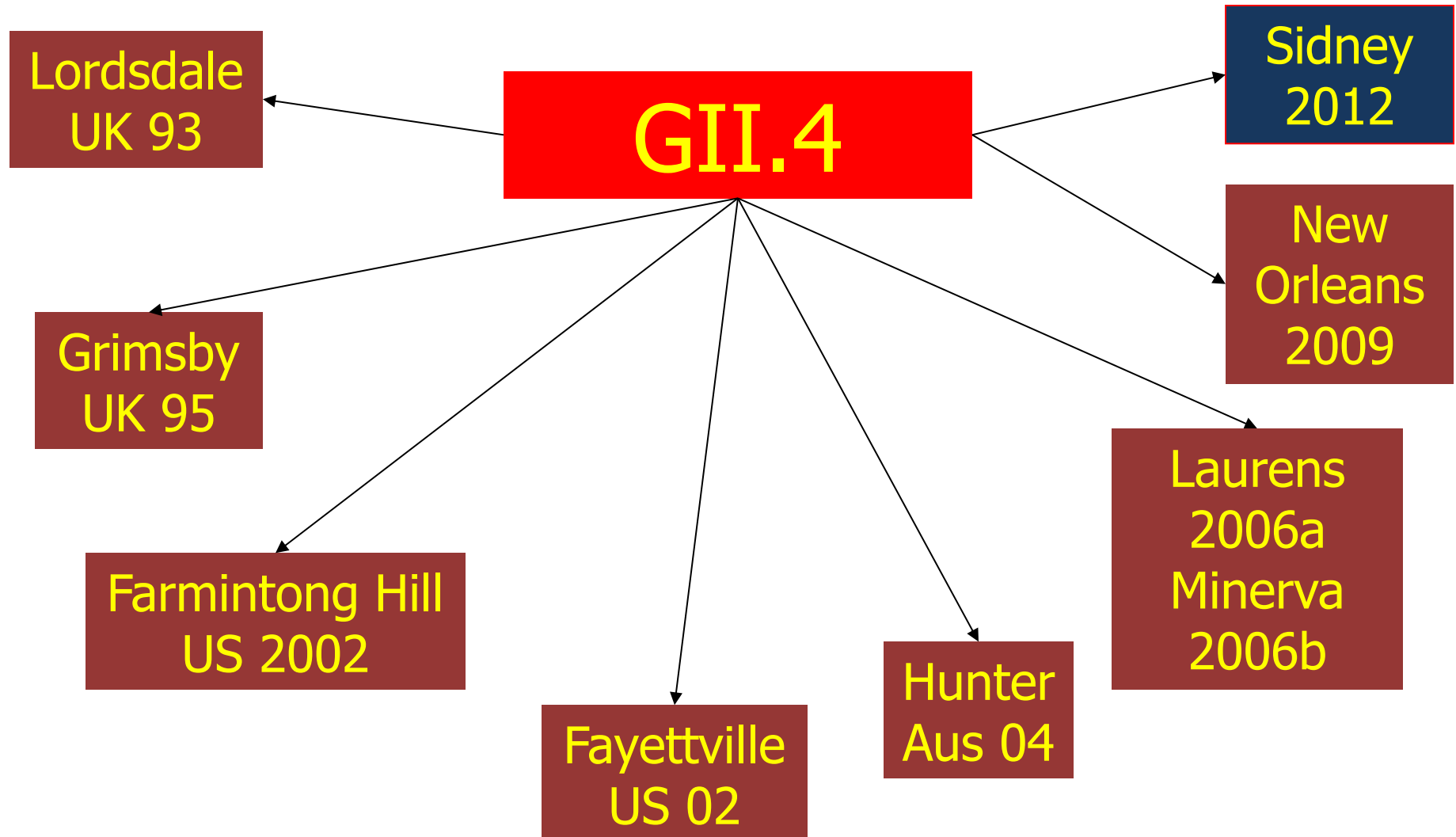
Genogroups



Genotypes

GII.1 GII.2 GII.3 **GII.4** GII.5 GII.6 GII.7 GII.8 GII.9 GII... GII.18

Representatives Strains of GII.4 Genotype



Treatment

- No specific treatment
- Fluids replacement recommended
- Antiemetic
- VLPs Vaccine?

Norovirus vaccine development: next steps

Expert Rev. Vaccines 11(9), 1023–1025 (2012)



Robert L Atmar

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Mary K Estes

*Departments of Medicine
and Molecular Virology &
Microbiology, Baylor College of
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USA*

“Although the disease is often self limited and recovery is the rule, norovirus infection is estimated to cause up to 10% of hospitalizations for gastroenteritis in the USA, up to 200,000 deaths in children under 5 years of age in developing countries, and mortality in the elderly.”

Noroviruses (NoVs) are a common cause of acute gastroenteritis in humans, with an estimated 21 million cases occurring in the USA annually [1]. Although the disease is often self limited and recovery is the rule, NoV infection is estimated to cause up to 10% of hospitalizations for gastroenteritis in the USA [2], up to 200,000 deaths in children under 5 years of age in developing countries [3], and mortality in the elderly

NoVs, in that failure to express certain HBGAs owing to a nonfunctional glycosylase has been associated with absolute resistance to infection (e.g., Norwalk virus and fucosyl transferase 2 [7]). HBGAs have been proposed to be an attachment factor or receptor for these NoVs. Adaptive immunity also influences susceptibility to infection in that the presence of serum antibodies that bind to VLPs and then

143

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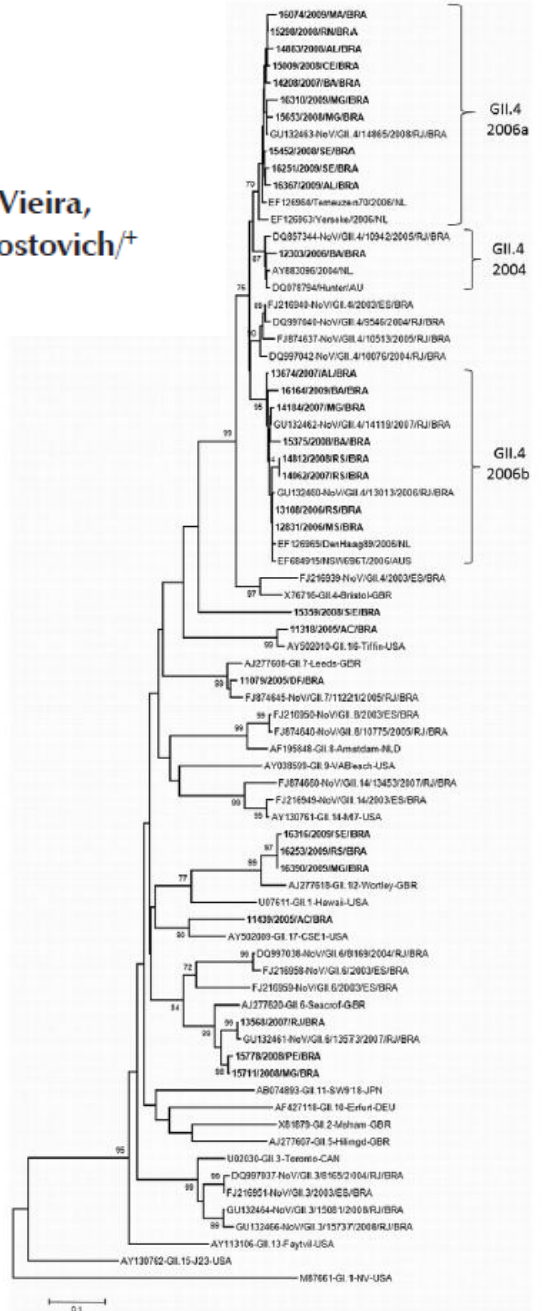
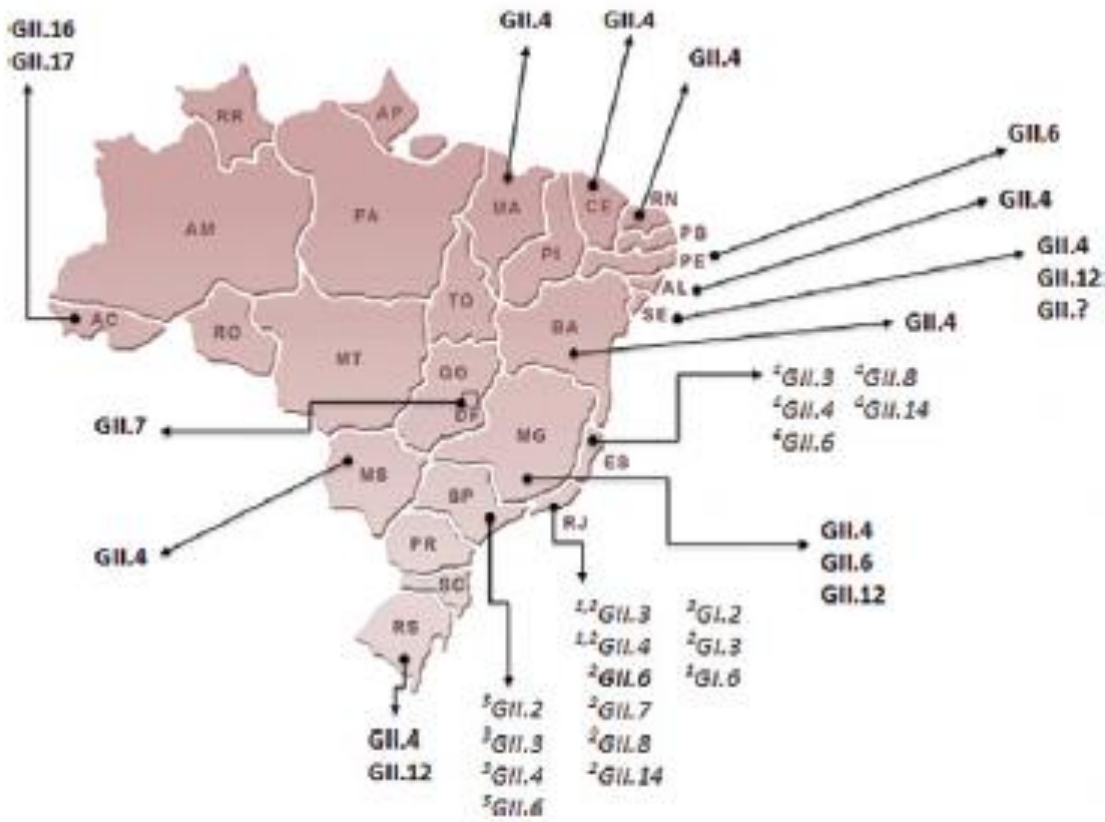
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Genetic diversity of noroviruses in Brazil

Julia Monassa Fioretti, Mônica Simões Rocha Ferreira, Matias Victoria, Carmen Baur Vieira, Maria da Penha Trindade Pinheiro Xavier, José Paulo Gagliardi Leite, Marize Pereira Miagostovich/+

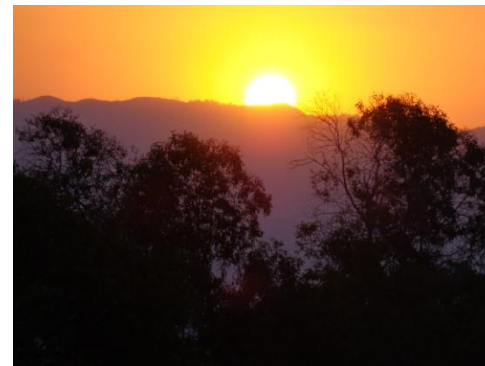
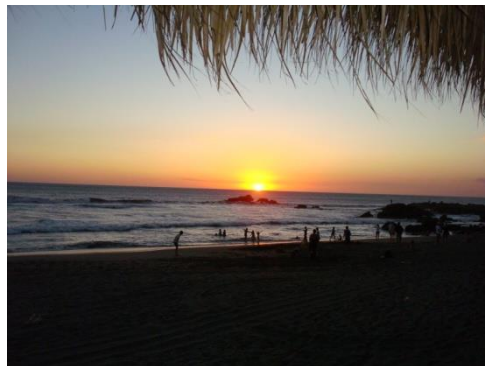
Laboratório de Virologia Comparada e Ambiental, Instituto Oswaldo Cruz-Fiocruz, Rio de Janeiro, RJ, Brasil



"There is nothing permanent except change"

Heraclitus

Webster et al. Microbiol. Rev., 56: 152 - 179, 1992




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 terça-feira, 15 de janeiro de 2008, 08:01 | **Online**

0 comentário(s)

Teclado infectado causa ga

Doença altamente contagiosa é provocada por

BBC Brasil - BBC

 - Um teclado de computador e um mouse inf
 que atingiu 103 pessoas em uma escola prim

isto é noticia!

Quarta-feira, 16 de Janeiro de 2008

Teclado infectado provoca surto de gastroenterite

Os utilizadores de computador não estão expostos apenas aos vírus informáticos. Prova disso, aconteceu numa escola em Washington, nos EUA, onde 103 pessoas, que partilhavam o aparelho, foram contaminados pelo vírus responsável pela gastroenterite.



Segundo um relatório divulgado recentemente pelo Centro norte-

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Concorda com a nova lei do Tabaco?
☐ Sim


Group A Rotavirus detection on environmental surfaces in hospital ICU ward.

Ana Carolina Ganime, MSc¹; Filipe A Carvalho-Costa, MD, PhD²; Marcos Cesar L Mendonça, PhD¹; Carmen B VieiraMSc¹; Marisa Santos, PhD³; Rubens Costa Filho, MD, MBA⁴; Marize P Miagostovich, PhD¹; José Paulo G Leite, PhD^{1*}

Table I: Monthly detection of group A rotavirus by nested-RT-PCR from environmental surface samples of an Intensive care unit in a hospital in Rio de Janeiro.

Fomites ^a	Months of year 2009						Total + /fomite ^b
	January	February	March ^c	April	May	June	
Keyboard infusion pumps	+ ^{2R}	+ ^{1R}	+ ^{1R}	-	-	-	4 (9%)
Interior door handles	+ ^{3R}	+ ^{2R}	+ ^{1R}	-	-	-	6 (14%)
Telephones	+ ^{1R}	+ ^{1R}	+ ^{1R}	-	-	-	3 (7%)
Coffee support tables	+ ^{2R}	-	-	-	+ ^{1R}	-	3 (7%)
Exterior bathroom door handle	+ ^{2R}	+ ^{3R}	-	-	-	-	5 (12%)
Alcohol gel supports	+ ^{6R}	+ ^{4R}	-	-	-	+ ^{1R}	11 (26%)
Chlorhexidine® supports	+ ^{3R}	+ ^{5R}	-	-	-	-	8 (19%)
Common trash bin covers	+ ^{2R}	+ ^{5R}	-	-	-	+ ^{1R}	8 (19%)
TV remote controls	-	+ ^{1R}	-	-	+ ^{2R}	-	3 (7%)
Bed controls	+ ^{1R}	-	-	+ ^{1R}	+ ^{3R}	-	5 (12%)
Flushing buttons	+ ^{2R}	+ ^{1R}	+ ^{1R}	+ ^{2R}	+ ^{3R}	-	9 (21%)
Companion chairs	+ ^{2R}	+ ^{1R}	-	+ ^{4R}	+ ^{1R}	-	8 (19%)
Total + /month	26 (31%)	24 (29%)	4 (5%)	7 (8%)	10 (12%)	2 (2%)	73/504 (14%)

+ : positive samples; - : negative samples. Number R = number of positive rooms. ^a12 samples were collected from each of 7 rooms; ^b 42 samples were collected from each fomite; ^c Cleaning protocol changed .



Representação dos pontos de coleta. A) botão de descarga, B) controle remoto da TV, C) telefone, D) cadeira do acompanhante, E) suporte para álcool gel e clorexidina, F) maçaneta interna da porta do leito e maçaneta externa do banheiro, G) tampa da lixeira de resíduos comuns, H) controle da cama, I) teclado da bomba de infusão, J) mesa de apoio.

Detection of enteric viruses in recreational waters of an urban lagoon in the city of Rio de Janeiro, Brazil

**Carmen Baur Vieira¹, Anna Carolina de Oliveira Mendes²,
Flávia Ramos Guimarães¹, Tulio Machado Fumian¹, José Paulo Gagliardi Leite¹,
Ana Maria Coimbra Gaspar², Marize Pereira Miagostovich^{1/+}**

Mem Inst Oswaldo Cruz, Rio de Janeiro, Vol. 108(1): 77-83, February 2013

The efficiency of concentration methods used to detect enteric viruses in anaerobically digested sludge

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Water Research, Volume 47, Issue 8, 15 May 2013, Pages 2797-2810

Detection and quantification of classic and emerging viruses by skimmed-milk flocculation and PCR in river water from two geographical areas

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Food and Environmental Virology, September 2013, Volume 5, Issue 3, pp 144-149

Optimization of an Adsorption-Elution Method with a Negatively Charged Membrane to Recover Norovirus from Lettuce

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CAPÍTULO 4

Colaboração entre sociologia e virologia ambiental para a implementação de políticas públicas nacionais

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Promoção da relação saúde–saneamento–cidade por meio da Virologia Ambiental

CARLOS JOSÉ SALDANHA MACHADO

MARIZE PEREIRA MIAGOSTOVICH

JOSÉ PAULO GAGLIARDI LEITE

RODRIGO MACHADO VILANI

1. Introdução. 2. Constituição Federal, direito à saúde e visão holística da cidade. 3. Características da metodologia da Virologia Ambiental. 4. Estatuto da Cidade – Lei nº 10.257/2001. 5. Política Nacional de Meio Ambiente – Lei nº 6.938/81. 6. Política Nacional de Recursos Hídricos – Lei nº 9.433/97. 7. Política Federal de Saneamento Básico – Lei nº 11.445/2007. 8. Síntese das relações entre as políticas públicas analisadas e a Virologia Ambiental. 9. Reflexões, conclusões e recomendações.

World Day for Water

March 22nd, 1993

United Nations General Assembly



Iguazu Falls

Laboratory of Comparative and Environmental Virology





Ministério
da Saúde



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Ministry of
Health



Thank you and Welcome

Obrigado e Bem Vindos



Corcovado / Christ



Ipanema beach



Efficacies of Sodium Hypochlorite and Quaternary Ammonium Sanitizers for Reduction of Norovirus and Selected Bacteria during Ware-Washing Operations

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Cross-contamination of ready-to-eat (RTE) foods with pathogens on contaminated tableware and food preparation utensils is an important factor associated with foodborne illnesses. To prevent this, restaurants and food service establishments are required to achieve a minimum microbial reduction of 5 logs from these surfaces. This study evaluated the sanitization efficacies of ware-washing protocols (manual and mechanical) used in restaurants to clean tableware items. Ceramic plates, drinking glasses and stainless steel forks were used as the food contact surfaces. These were contaminated with cream cheese and reduced-fat milk inoculated with murine norovirus (MNV-1), *Escherichia coli* K-12 and *Listeria innocua*. The sanitizing solutions tested were sodium hypochlorite (chlorine), quaternary ammonium (QAC) and tap water (control). During the study, the survivability and response to the experimental conditions of the bacterial species was compared with that of MNV-1. The results showed that current ware-washing protocols used to remove bacteria from tableware items were not sufficient to achieve a 5 log reduction in MNV-1 titer. After washing, a maximum of 3 log reduction in the virus were obtained. It was concluded that MNV-1 appeared to be more resistant to both the washing process and the sanitizers when compared with *E. coli* K-12 and *L. innocua*.