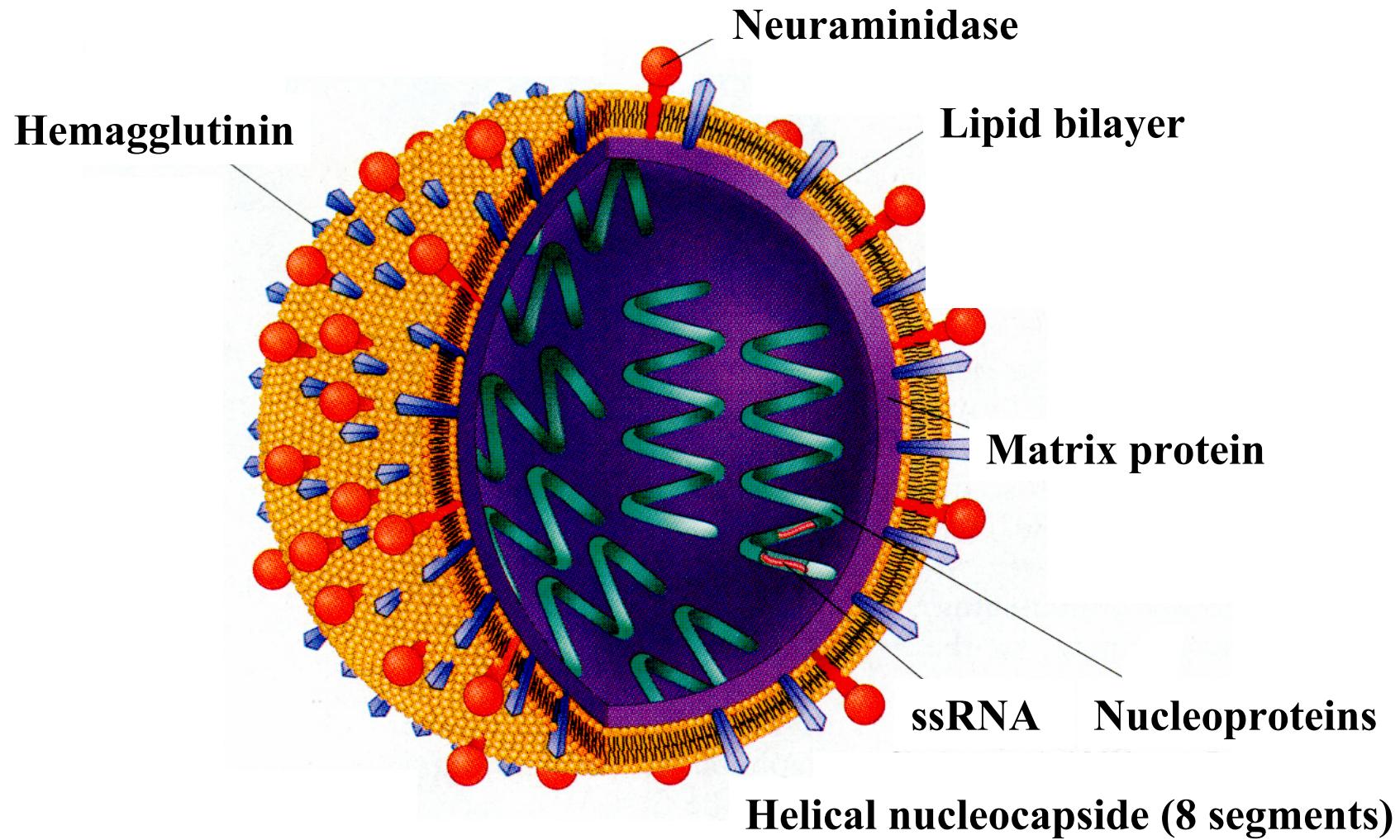


INFLUENZA A new pandemic ?

INFLUENZA VIRUS STRUCTURE



HEMAGGLUTININ SUBTYPES

Subtype	Human	Pig	Horse	Birds
H1	■			
H2	■			
H3	■			
H4				
H5	■			
H6				
H7	■			
H8				
H9	■			
H10				
H11				
H12				
H13				
H14				
H15				

NEURAMINIDASE SUBTYPES

Subtype	Human	Pig	Horse	Birds
N1	■			
N2	■			
N3				
N4				
N5				
N6				
N7	■			
N8				
N9			■	■■■■■■■■■■

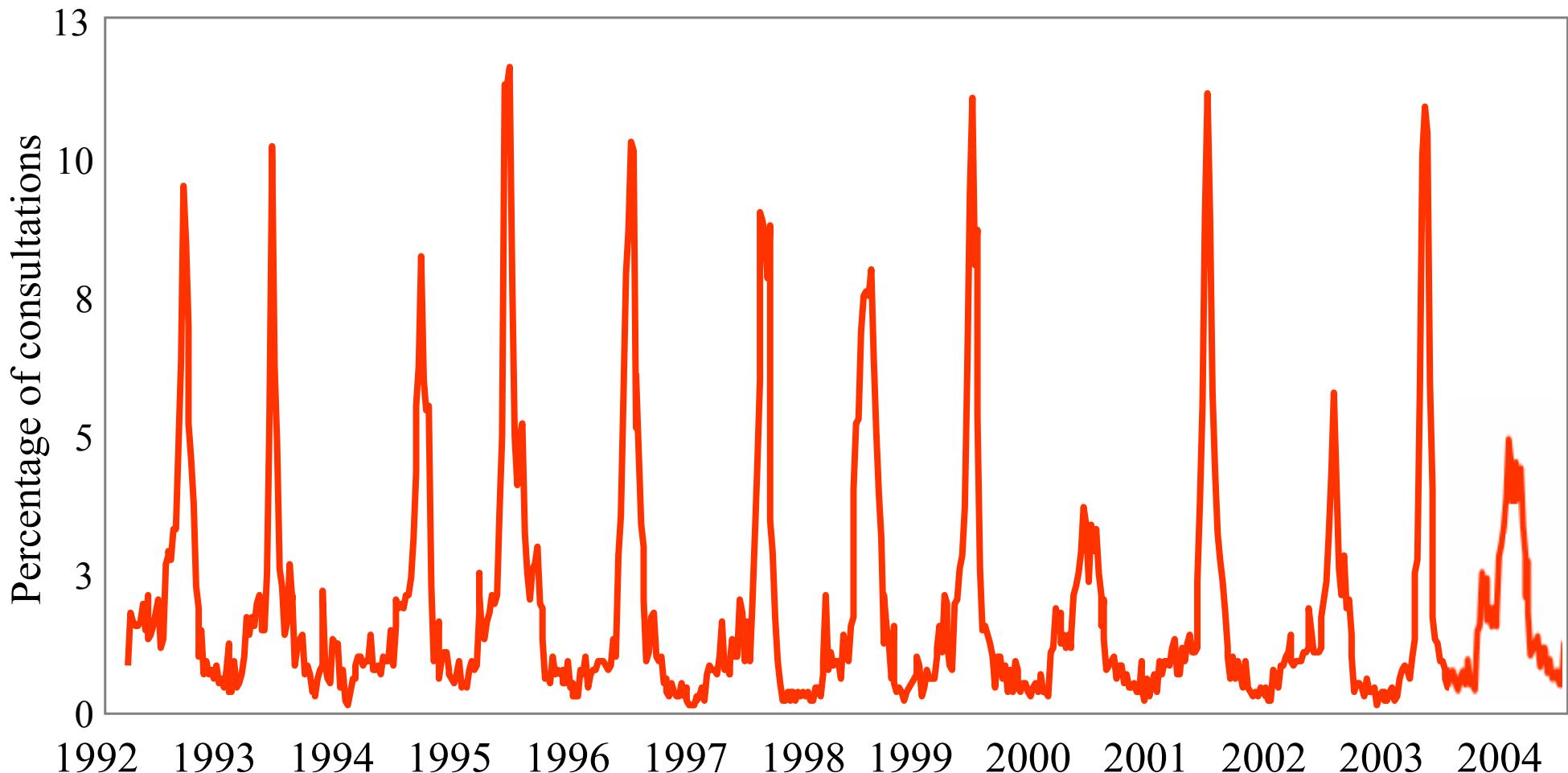
ANTIGENIC DRIFT

H3 → H3' → H3''

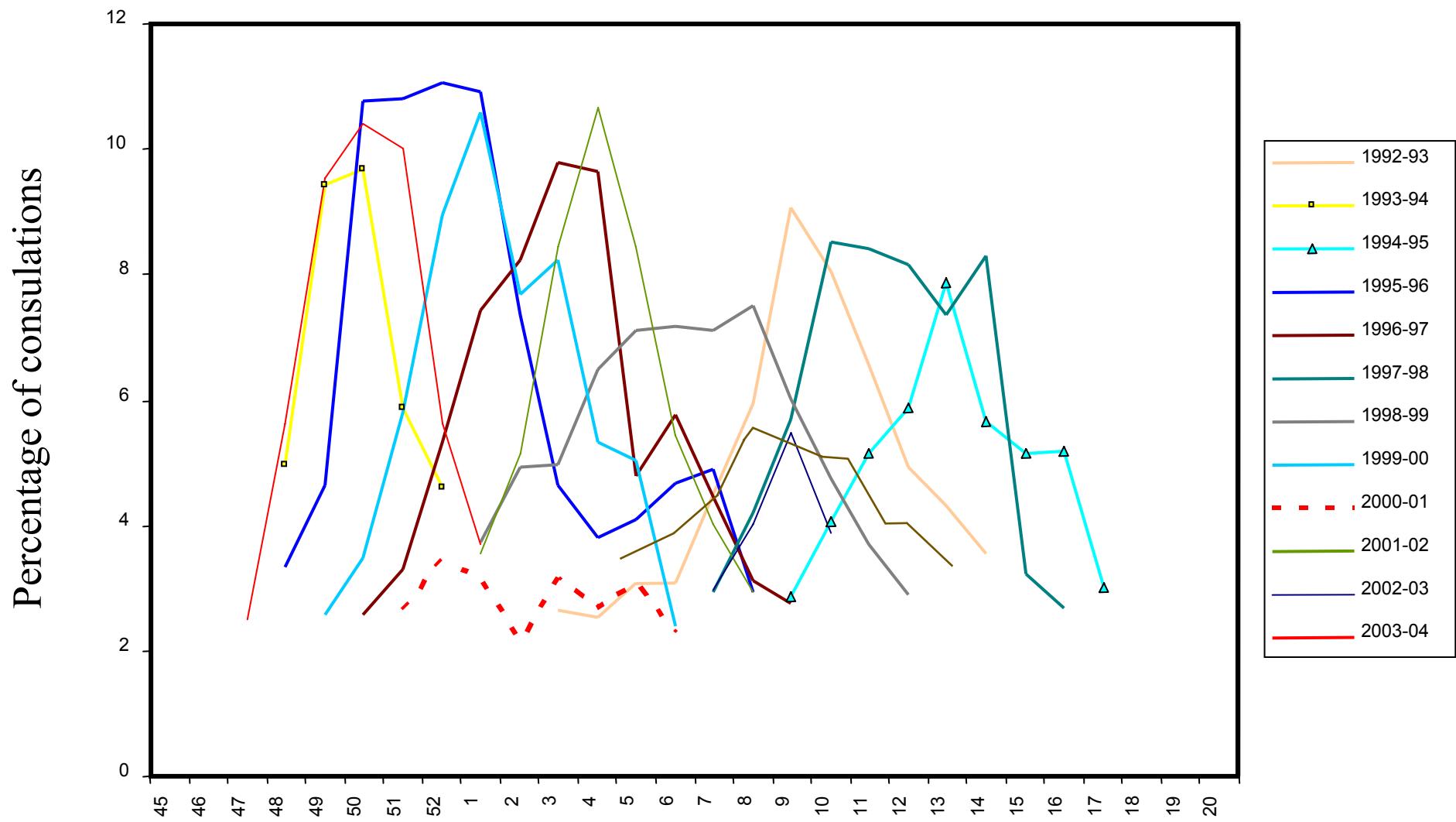
N2 → N2' → N2''



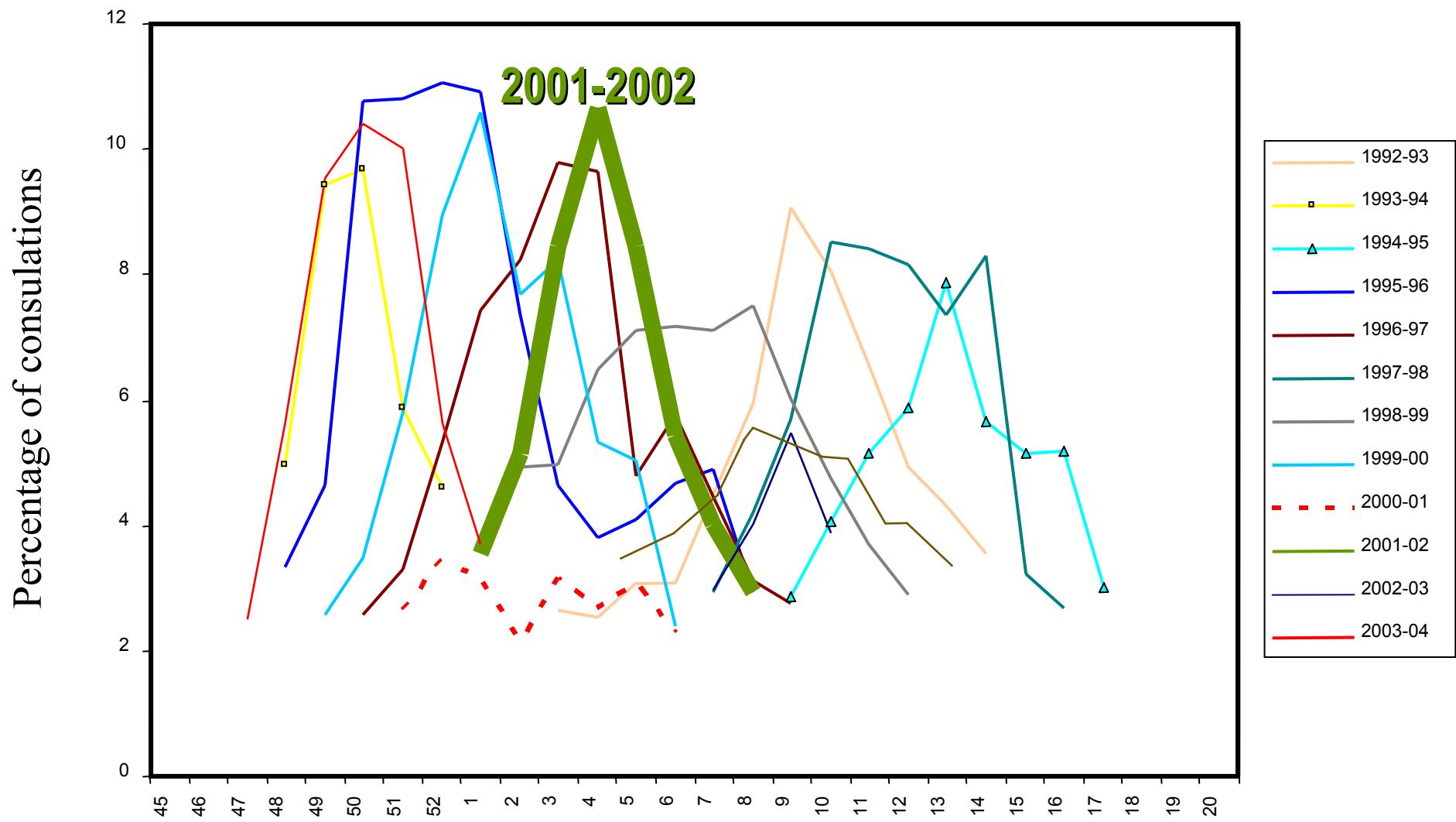
INFLUENZA SEASONALITY



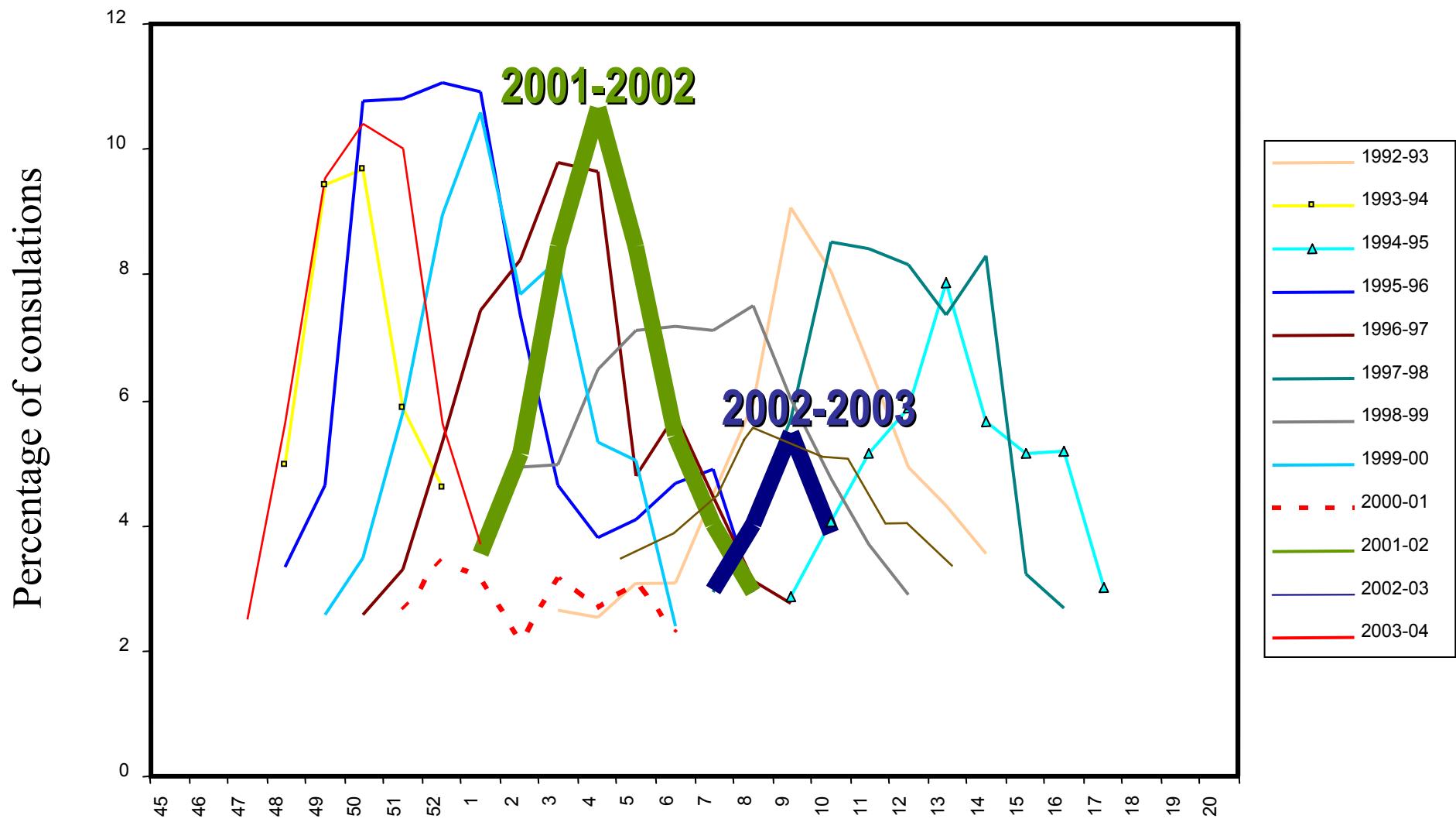
INFLUENZA 1992-2005



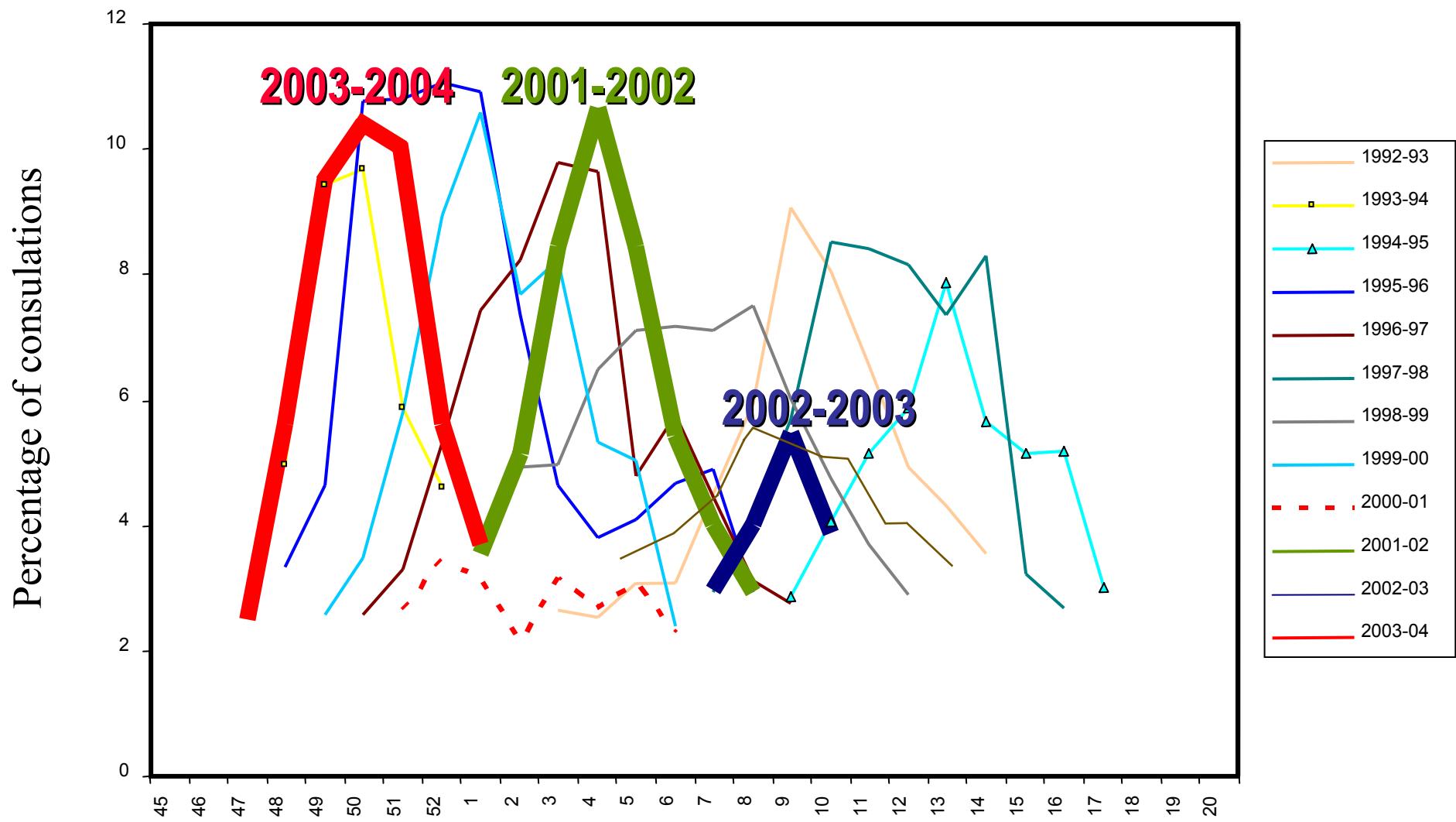
INFLUENZA 1992-2005



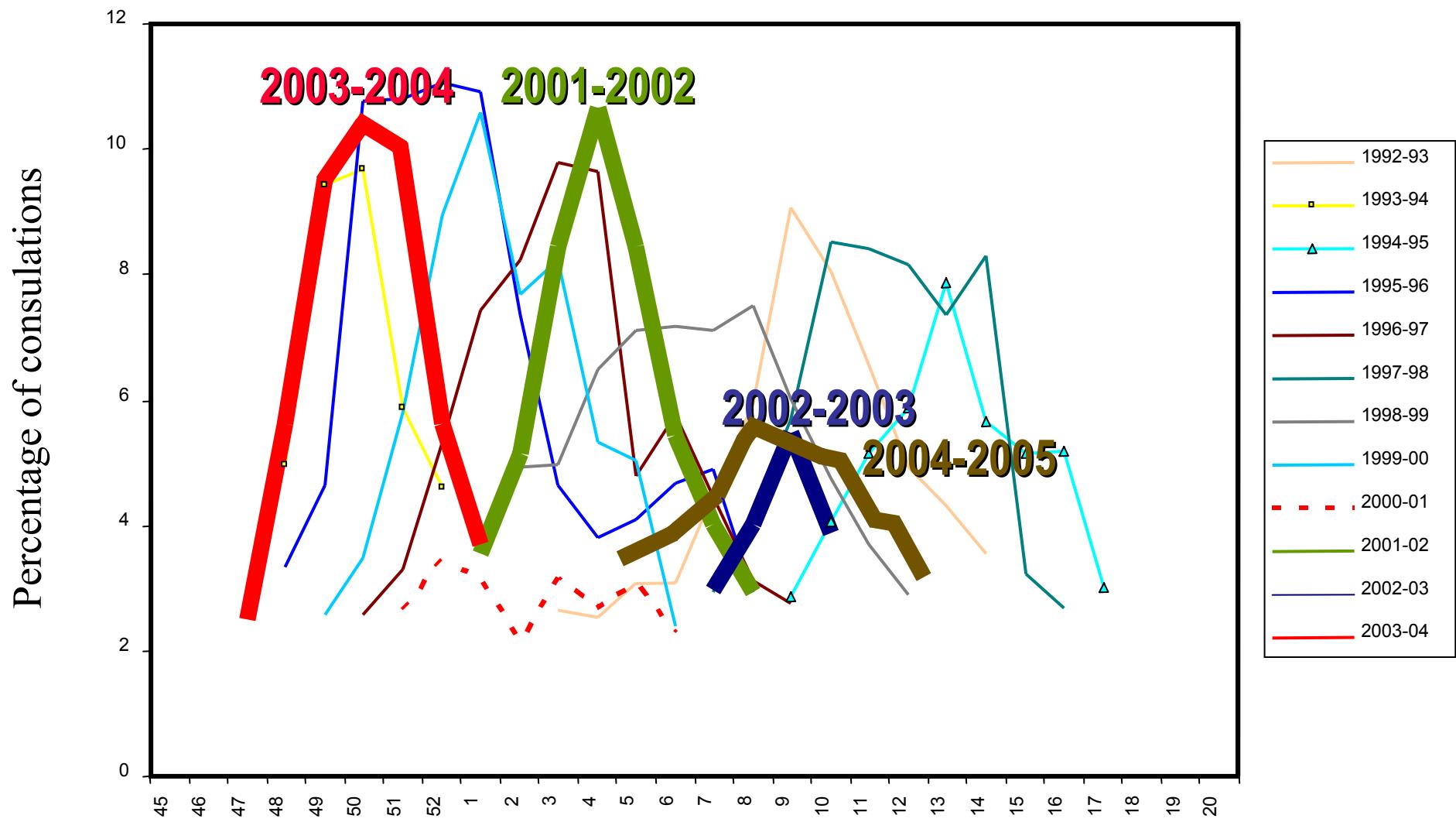
INFLUENZA 1992-2005



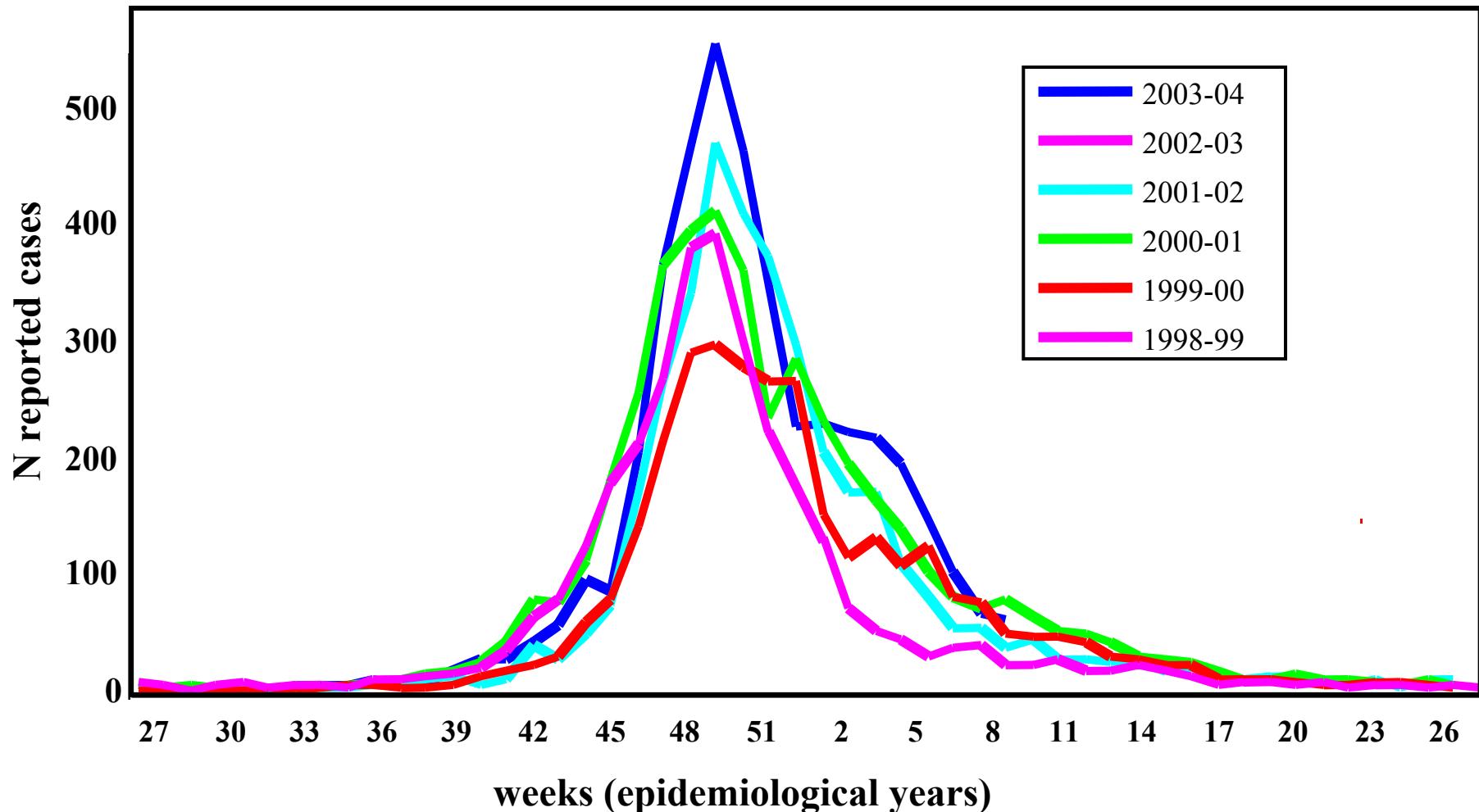
INFLUENZA 1992-2005



INFLUENZA 1992-2005

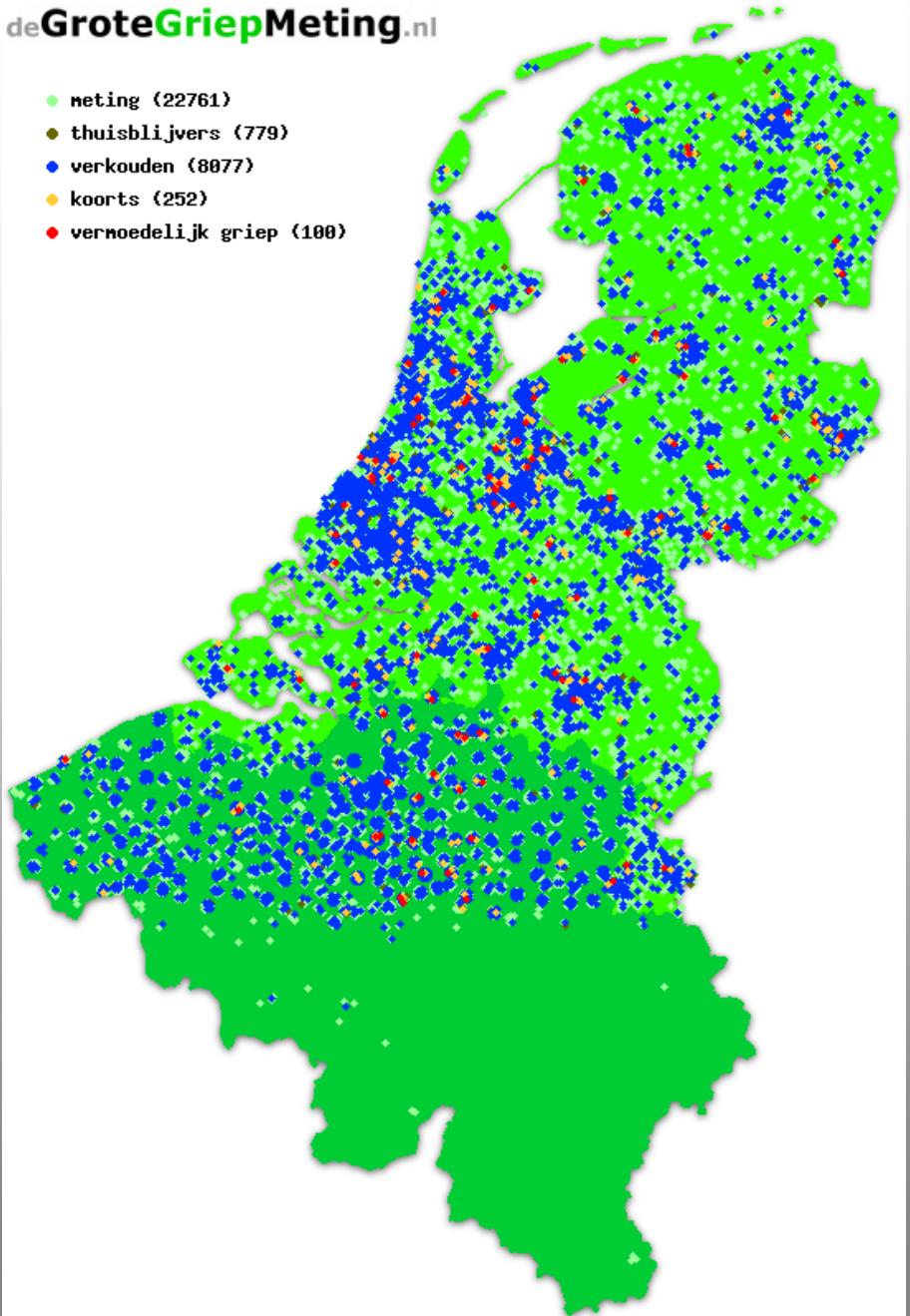


RSV INFECTIONS 1998-2004



deGroteGriepMetting.nl

- meting (22761)
- thuisblijvers (779)
- verkouden (8877)
- koorts (252)
- vermoedelijk griep (100)



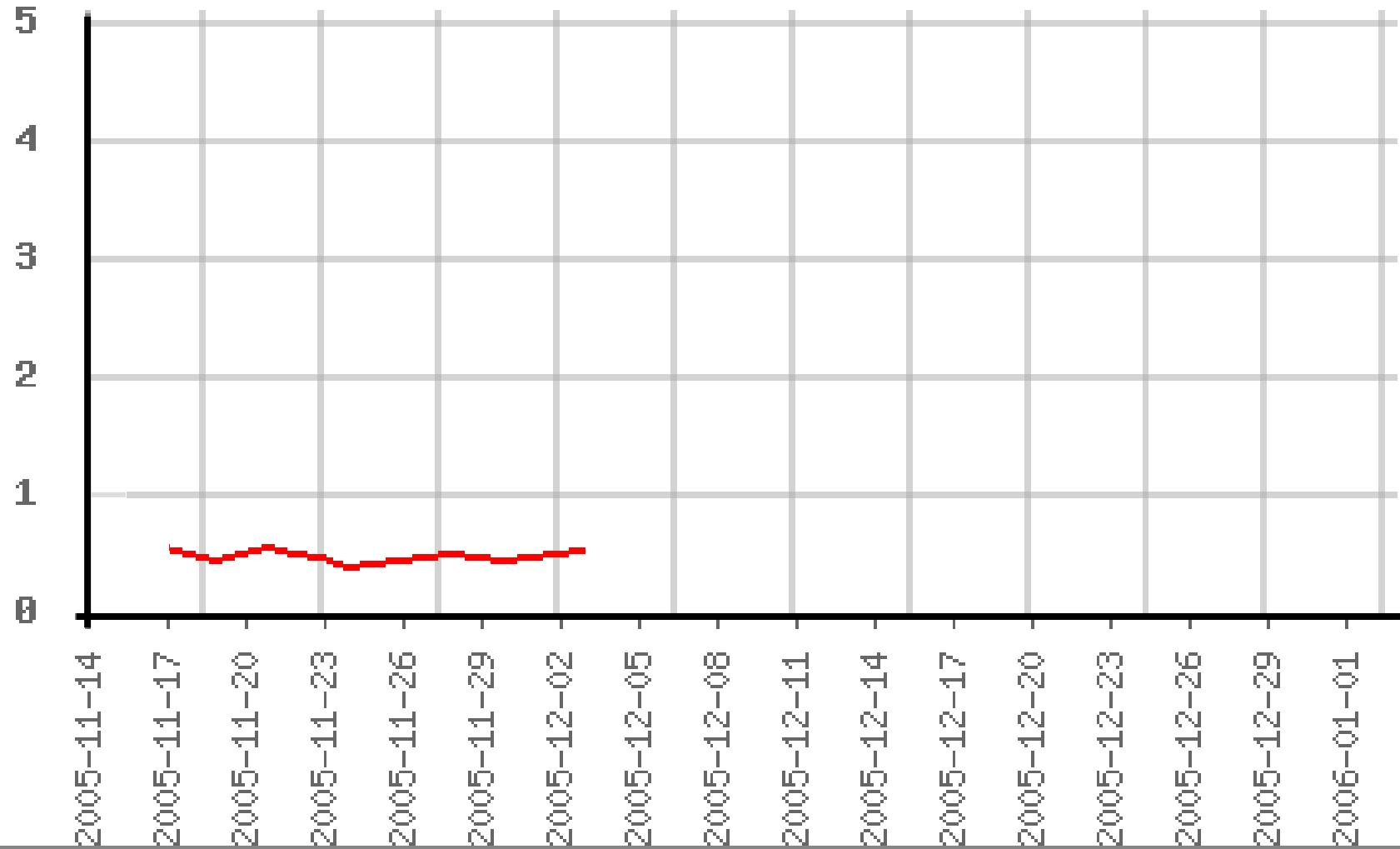
deGroteGriepMetting.nl

- vermoedelijk griep (48)



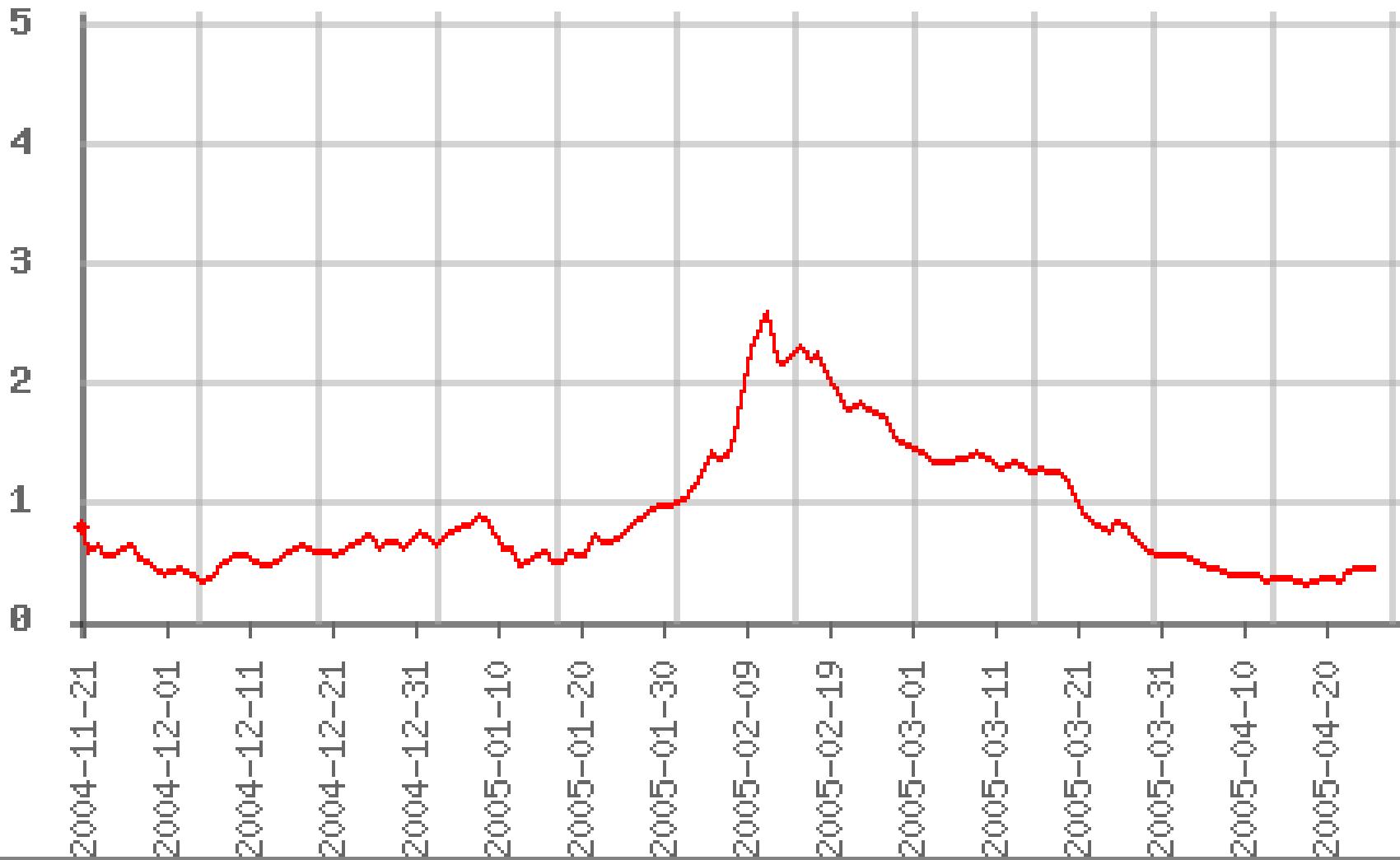
deGroteGriepMeting.nl

◆ percentage griep (gem. 0.53%)

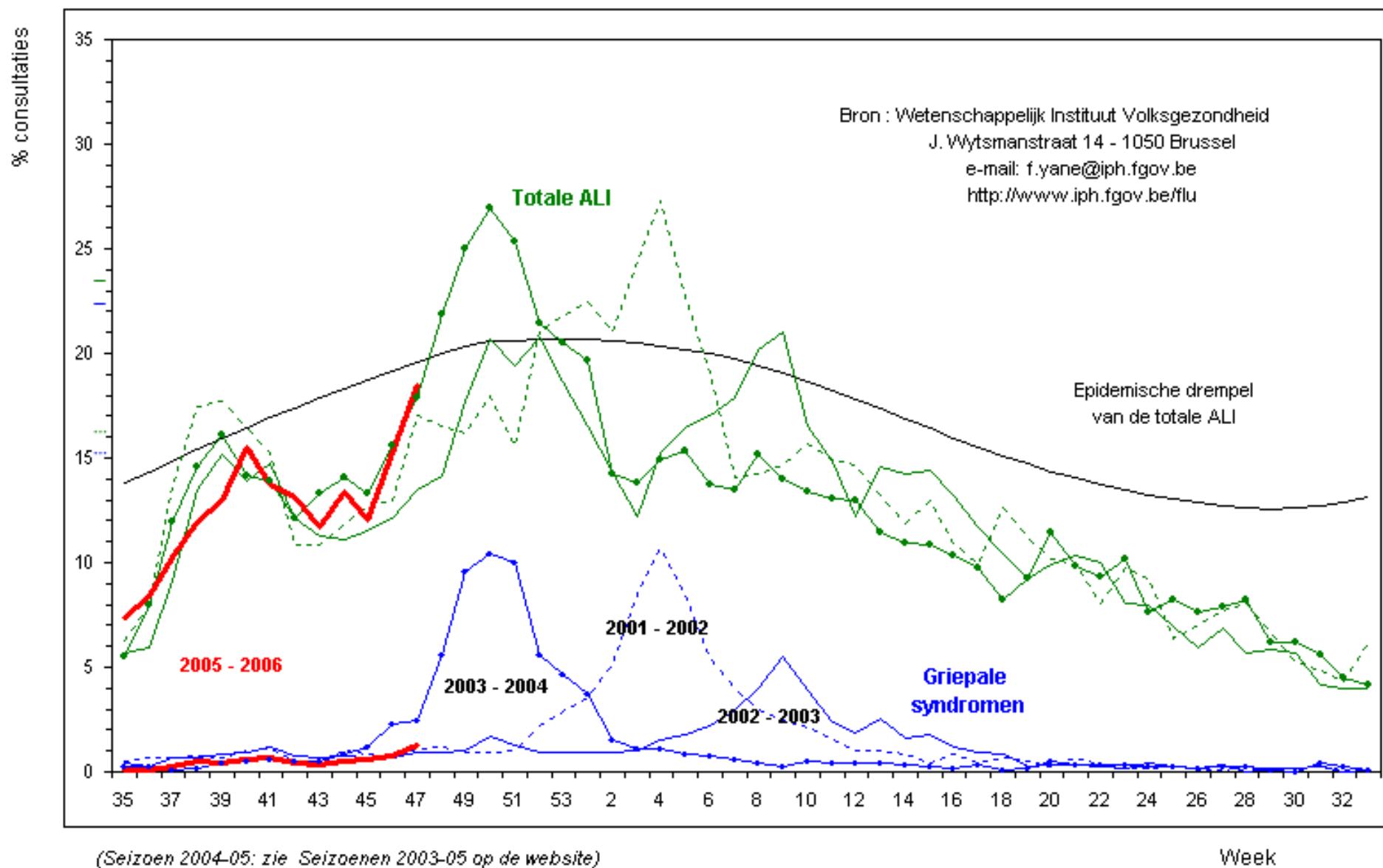


deGroteGriepMeting.nl

percentage griep (gem. 0.88%)



Evolutie van het percentage griepgevallen en acute luchtwegeninfecties (ALI) bij patienten geregistreerd door huisartsenpraktijken



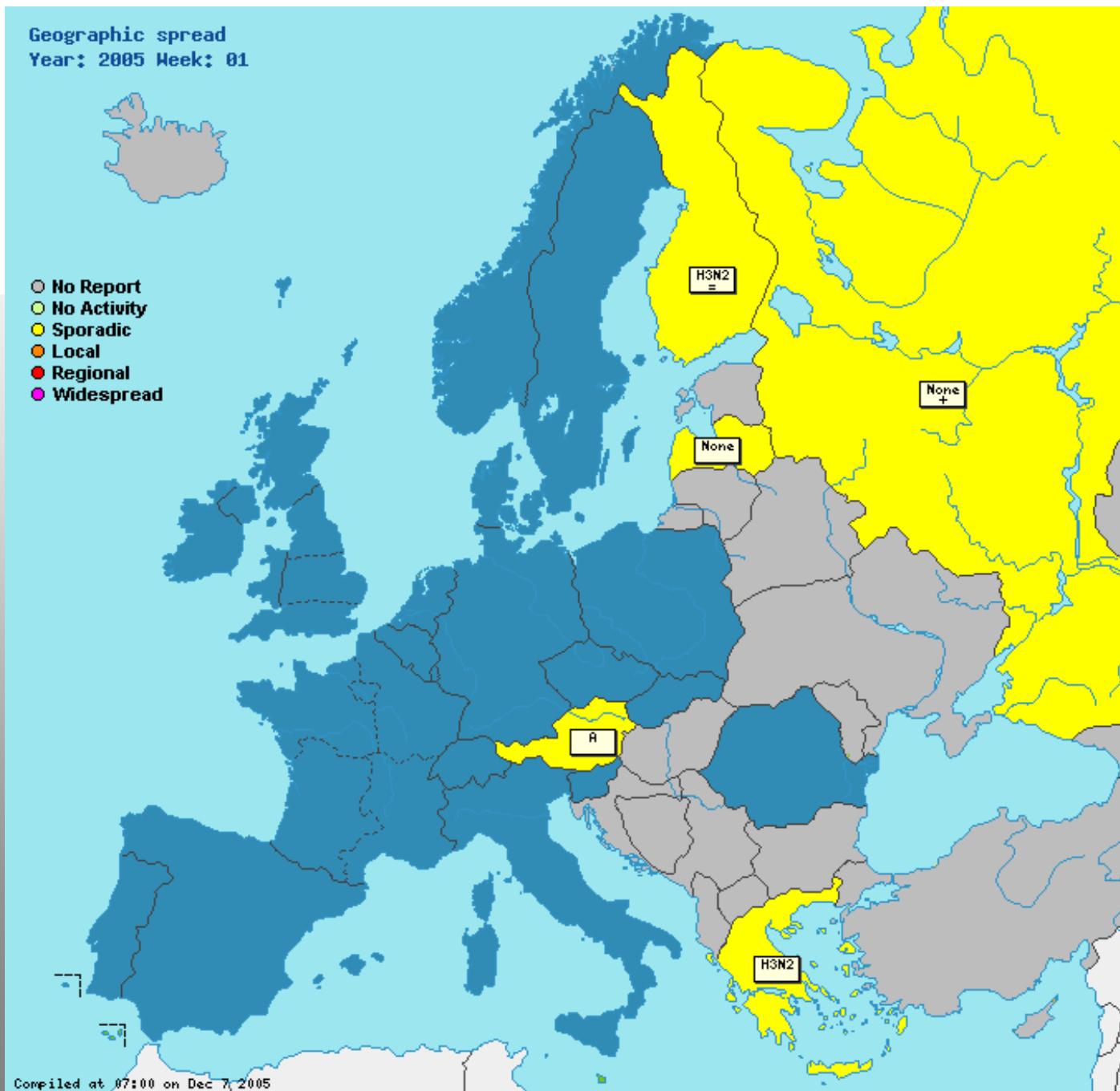


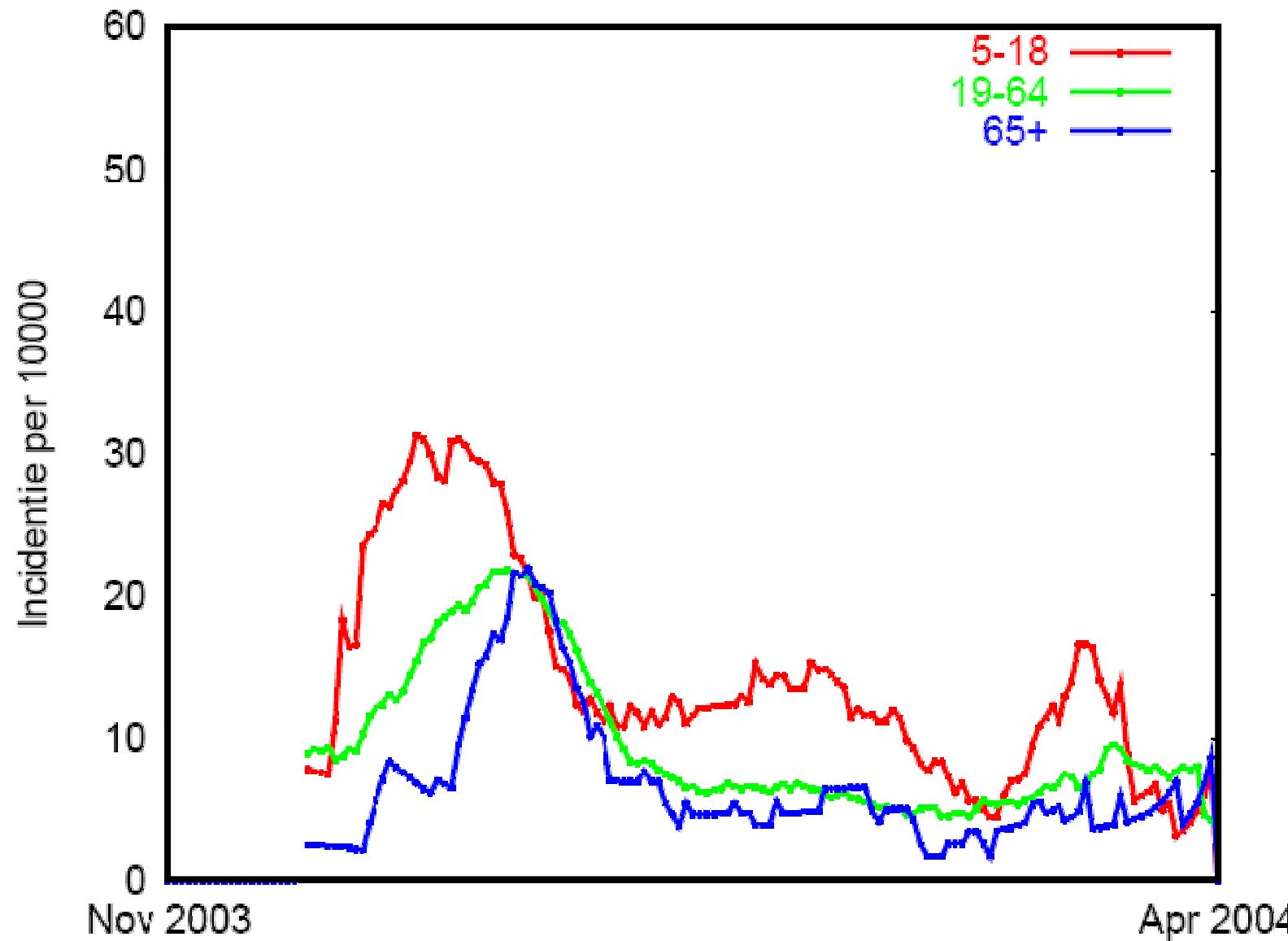


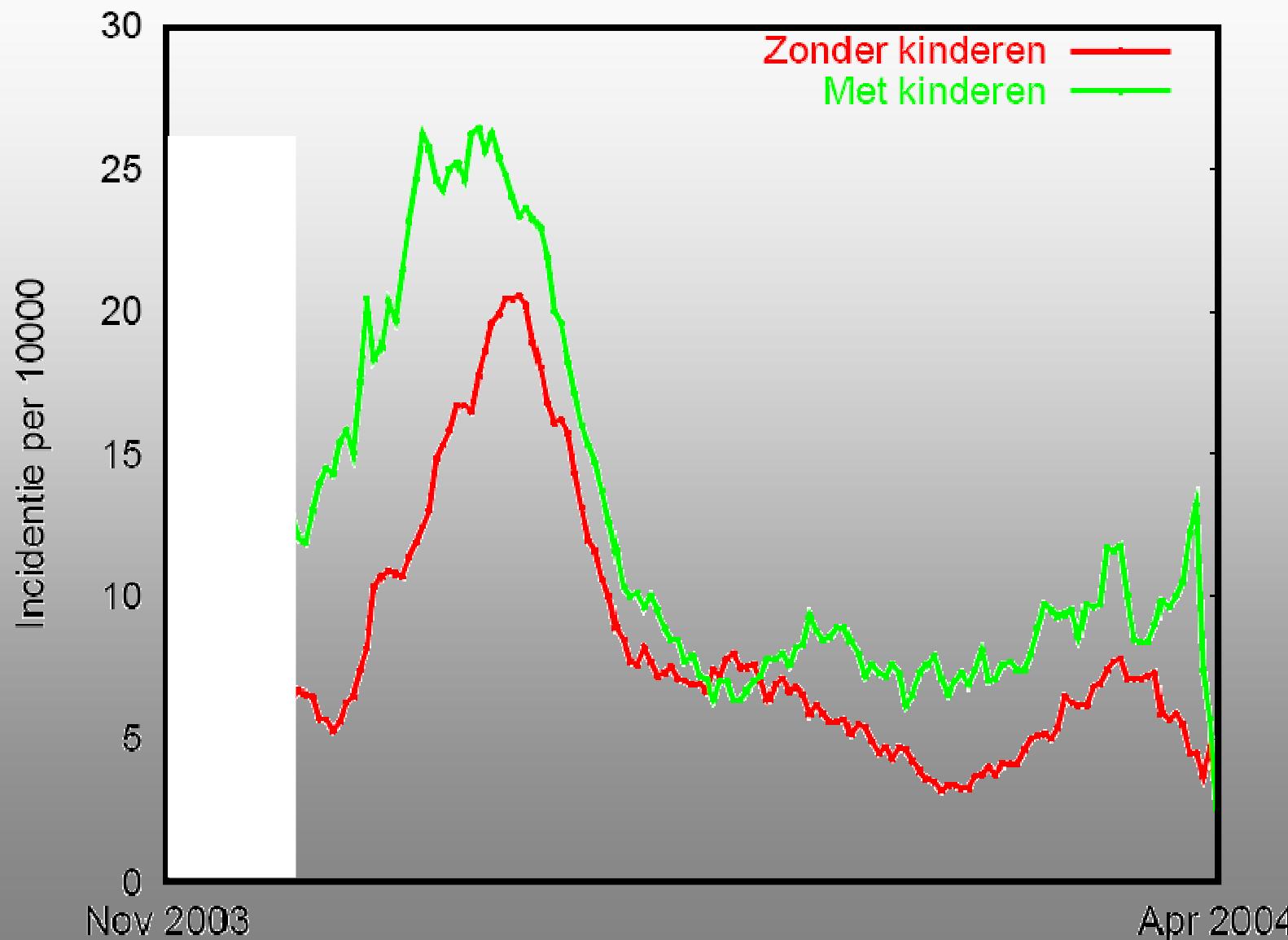
Geographic spread
Year: 2005 Week: 01



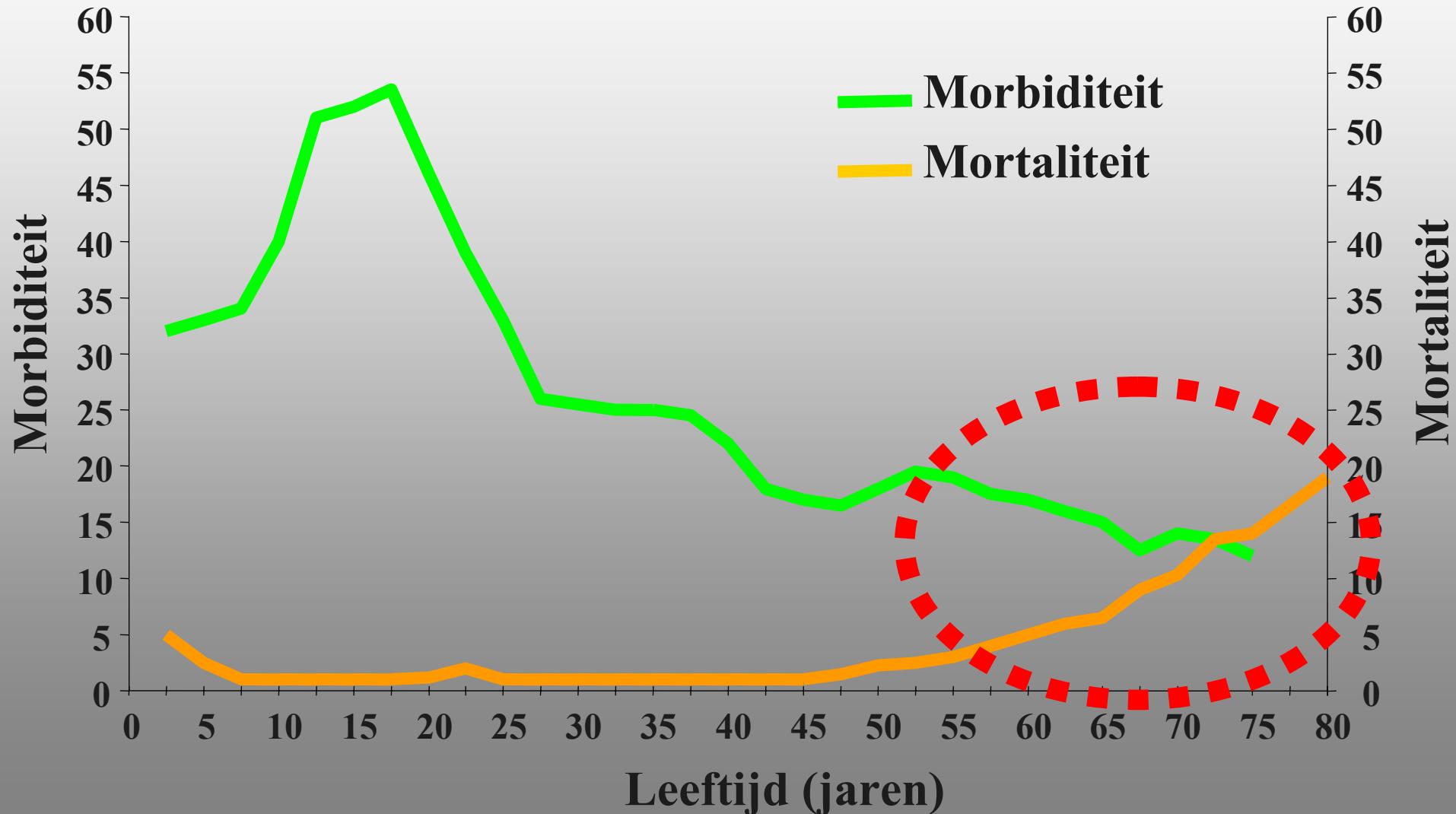
- No Report
- No Activity
- Sporadic
- Local
- Regional
- Widespread







INFLUENZA EPIDEMIOLOGIE





*Helping Families Honor the
Lives of Those They Love*



Hillebrand Industries, 1st Quarter Report, March 30, 2001

In the Funeral Services Group, Batesville Casket Company posted sales of \$163 million, a \$5 million decrease from the prior year. The company attributes the lower sales to a mild flu season resulting in a lower death rate and lower volume at Batesville Casket Company. Despite lower sales, gross margins increased to 50.9% in the current year, compared to 48.8% last year, primarily as a result of productivity improvements and improved product mix partially offset by lower volume.



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Vaccines, 'Wellcome' Brand—*continued*

'WELLCOME' BRAND—

„ Influenza Vaccine

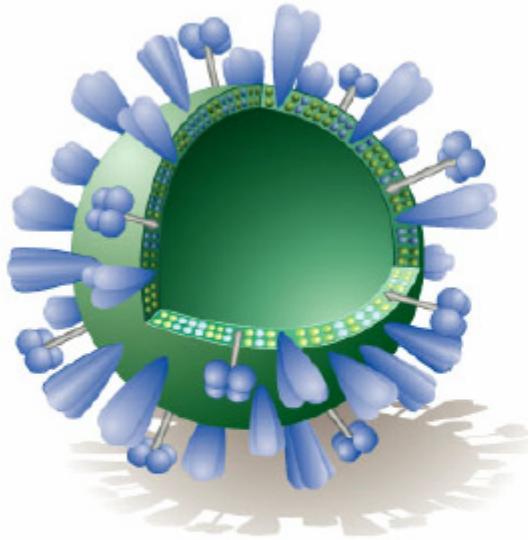
I c.c. containing 10 million *B. influenzae*

I c.c. „ 50 „ „ „





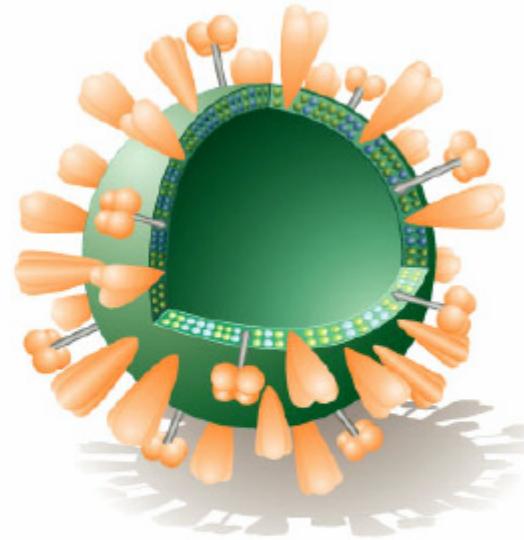
TRIVALENT INFLUENZA VACCINE



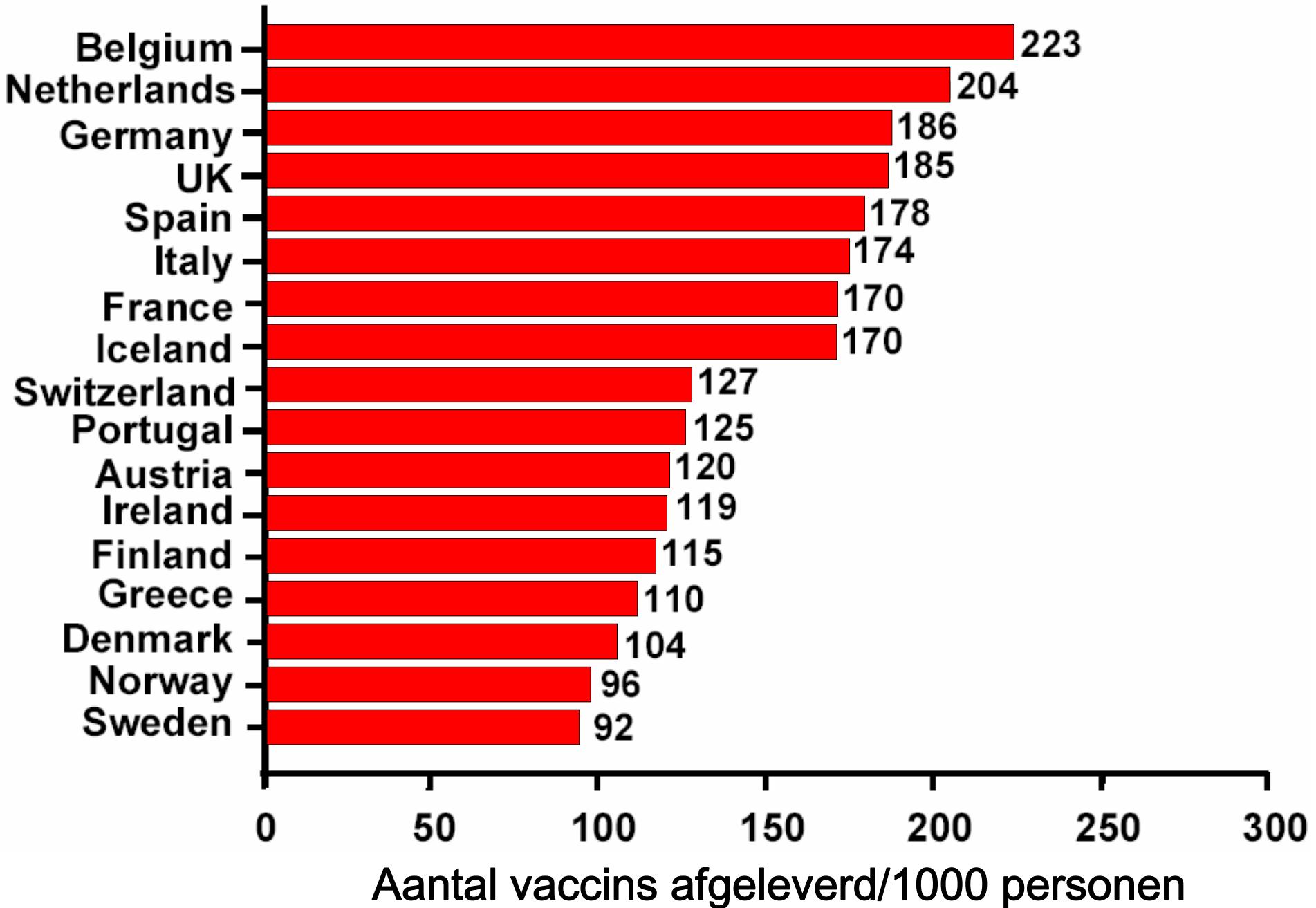
A/H1N1



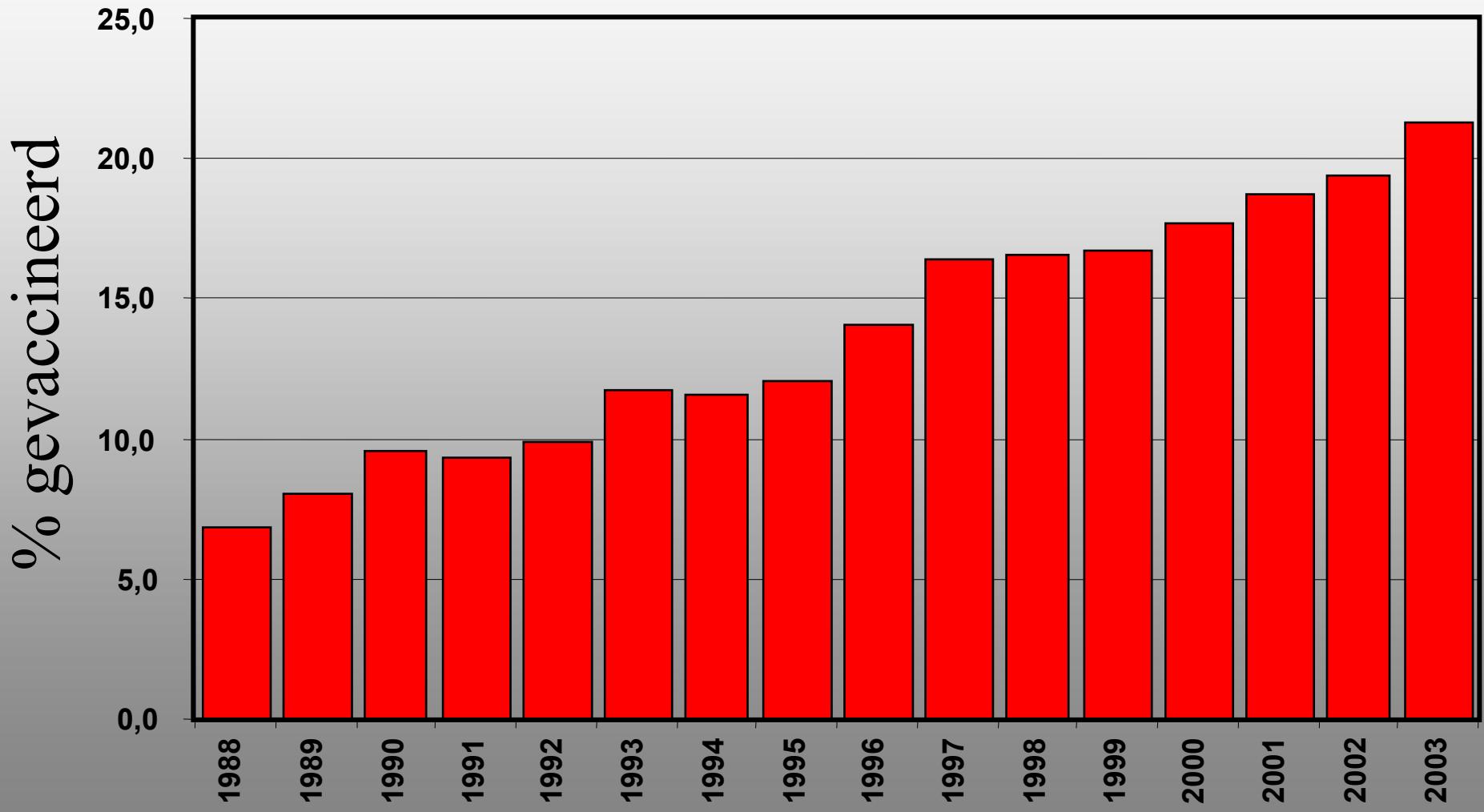
A/H3N2



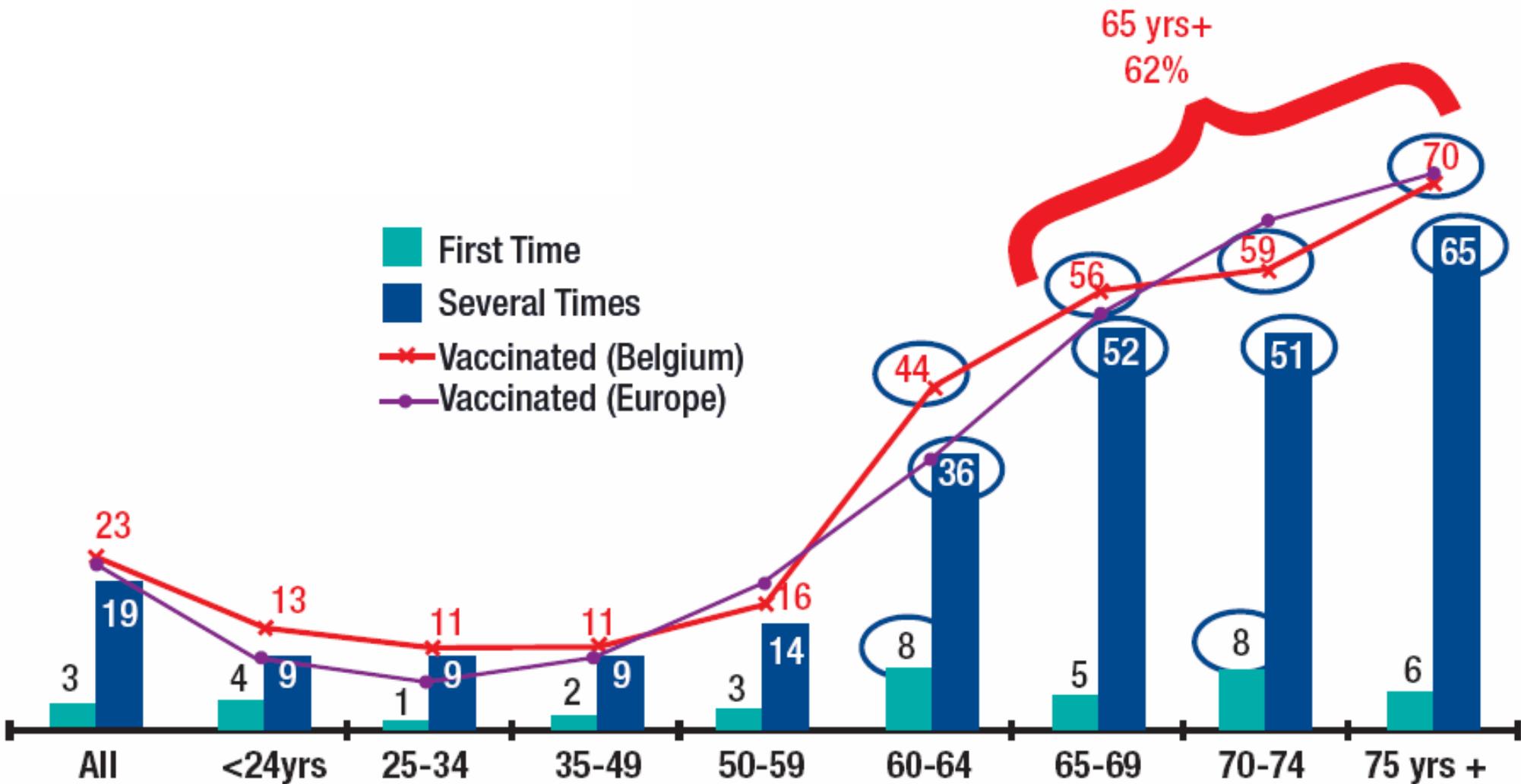
B



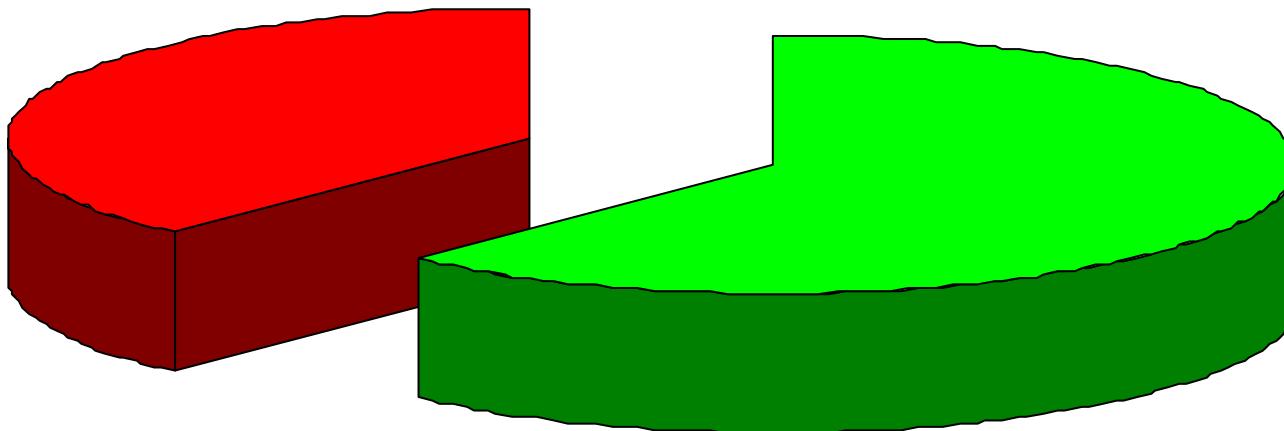
VACCINATION COVERAGE



VACCINATION COVERAGE



VACCINATION COVERAGE >65 year



Non-vaccinated

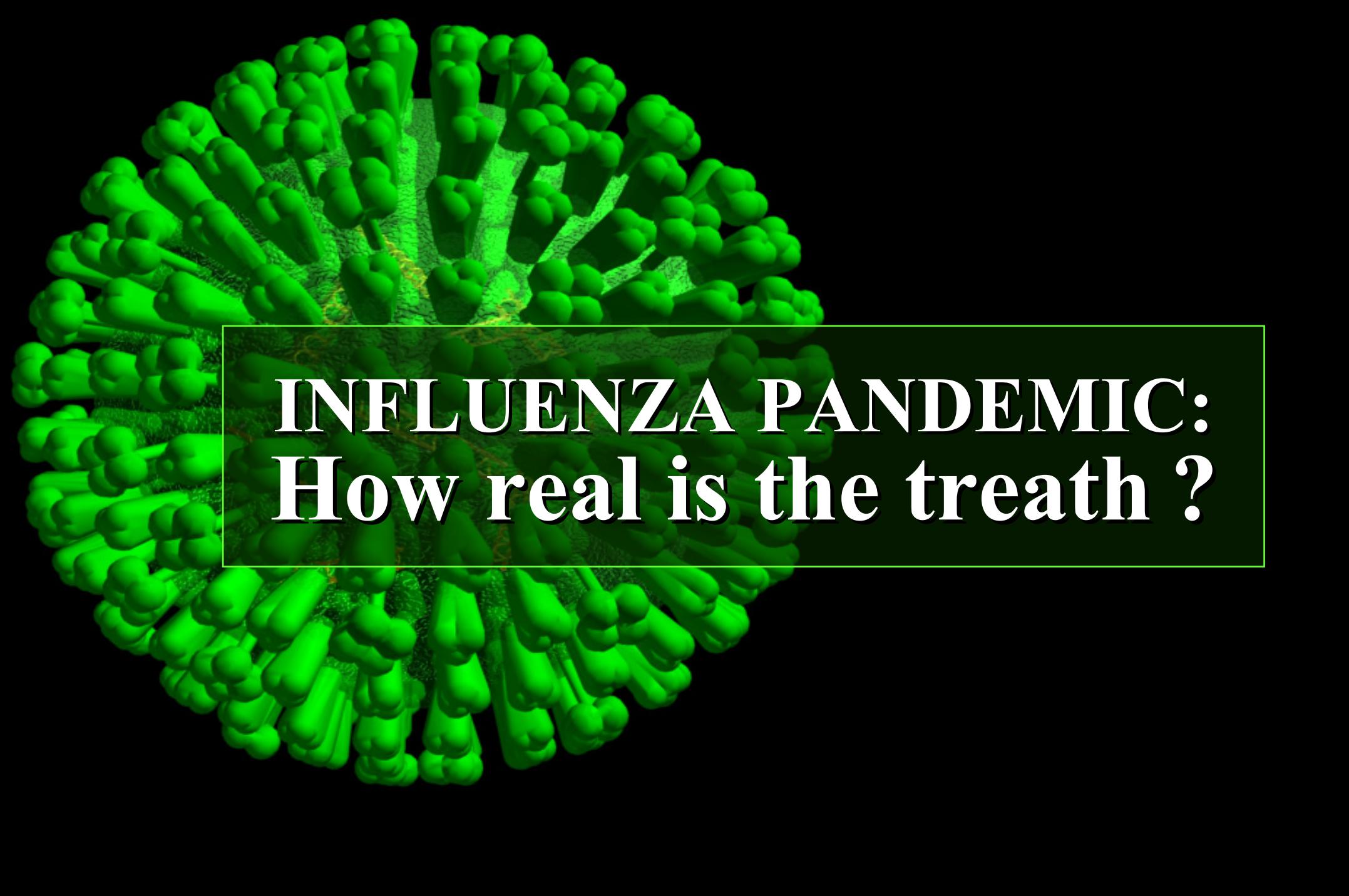
38%

700.000

Vaccinated

62%

1.140.000



INFLUENZA PANDEMIC: How real is the threat ?

GENETIC SHIFT

H3 → H5

N2 → N1



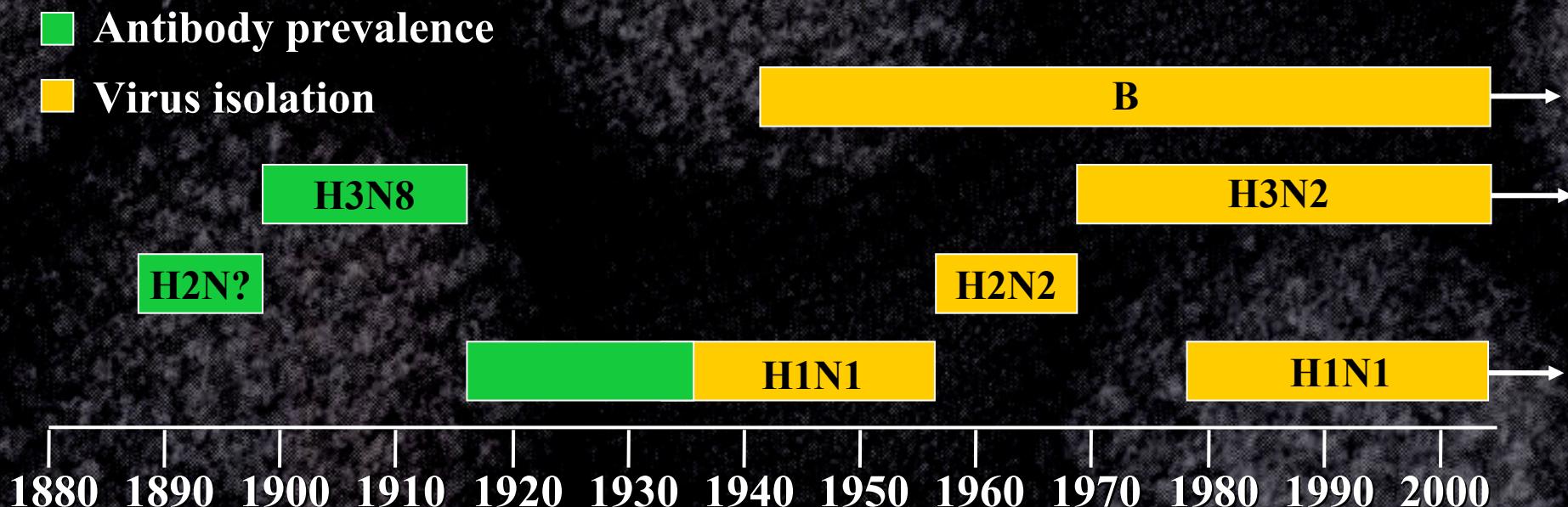
Year 1

Year 2

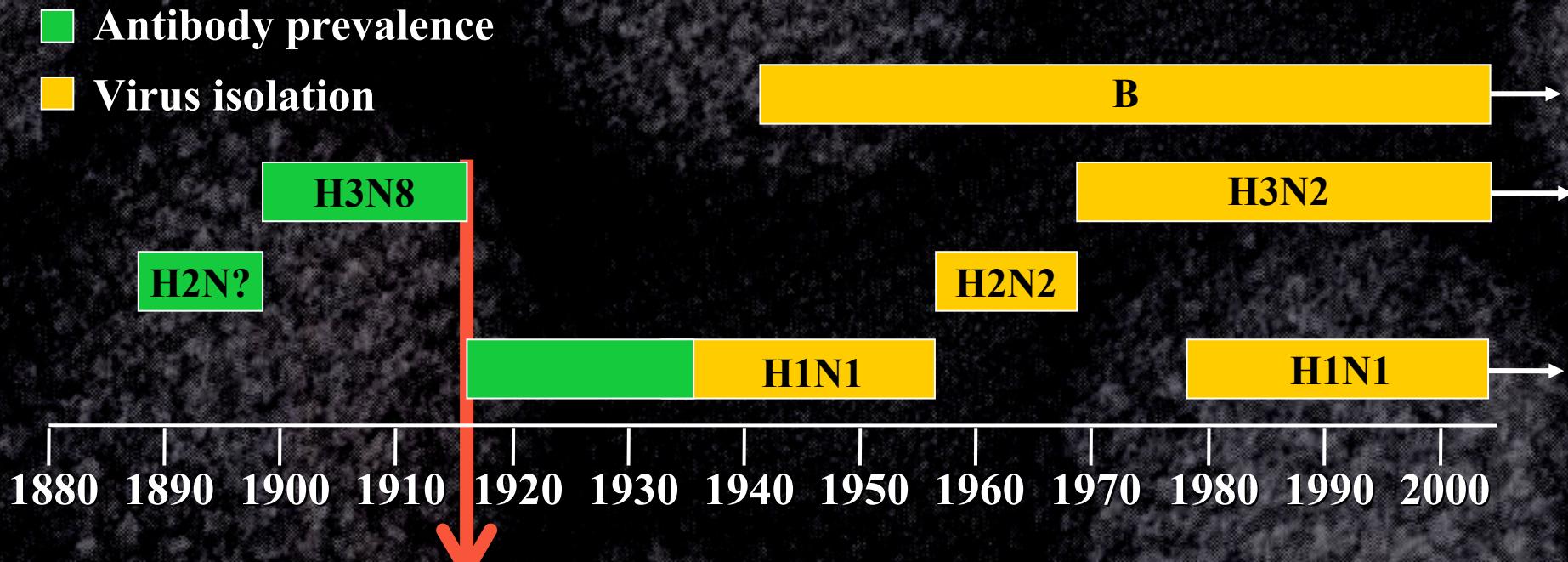


LA GRIPPE FAISANT LE TOUR DU MONDE.

INFLUENZA VIRUSES



INFLUENZA VIRUSES



1918 H1N1:
Spanish Flu



Kamp Funston, Kansas, 1918













Toronto Sky Dome



1918 SBIMC-BVIKM meeting

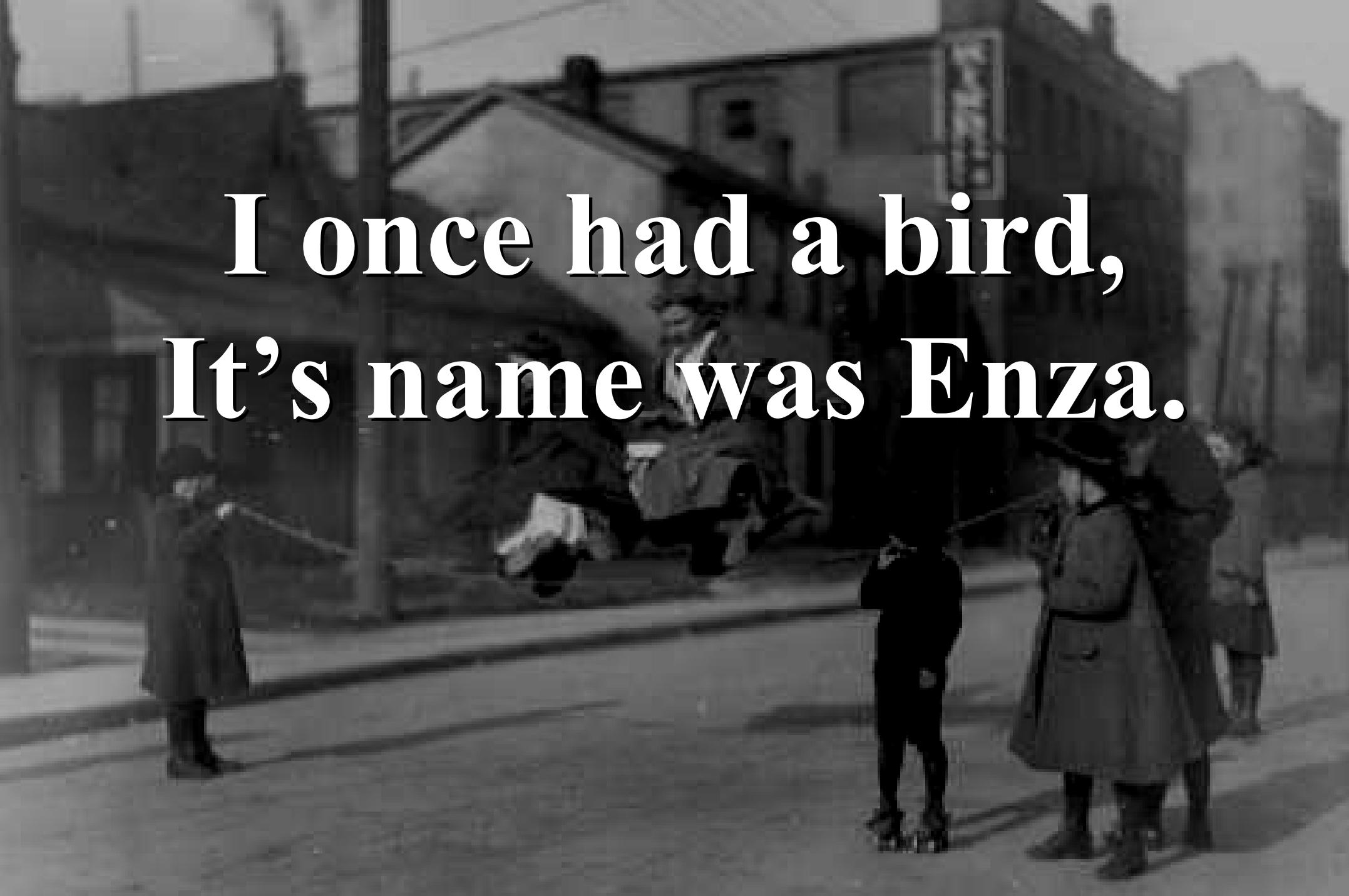


Hong Kong

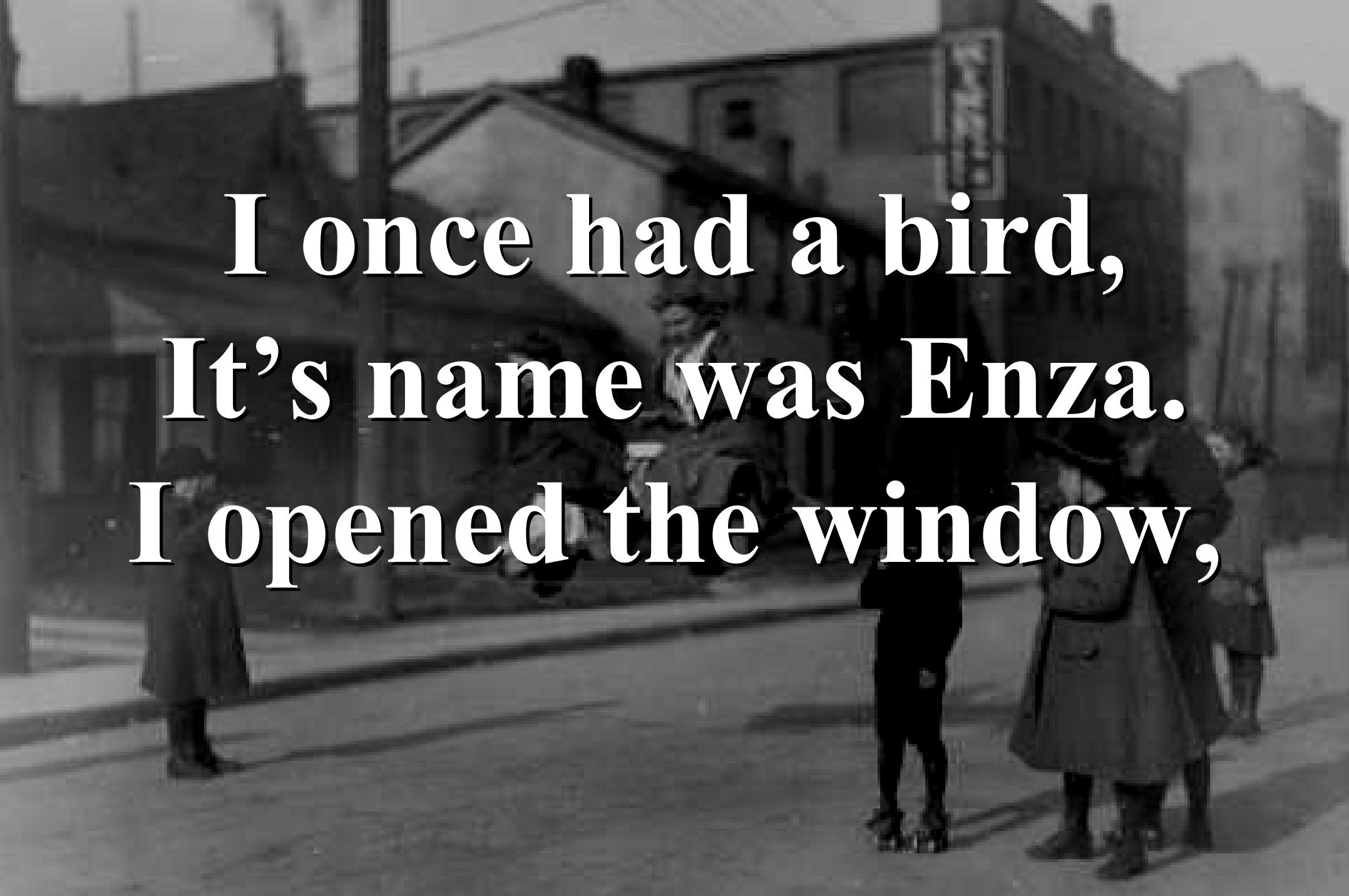


I once had a bird,

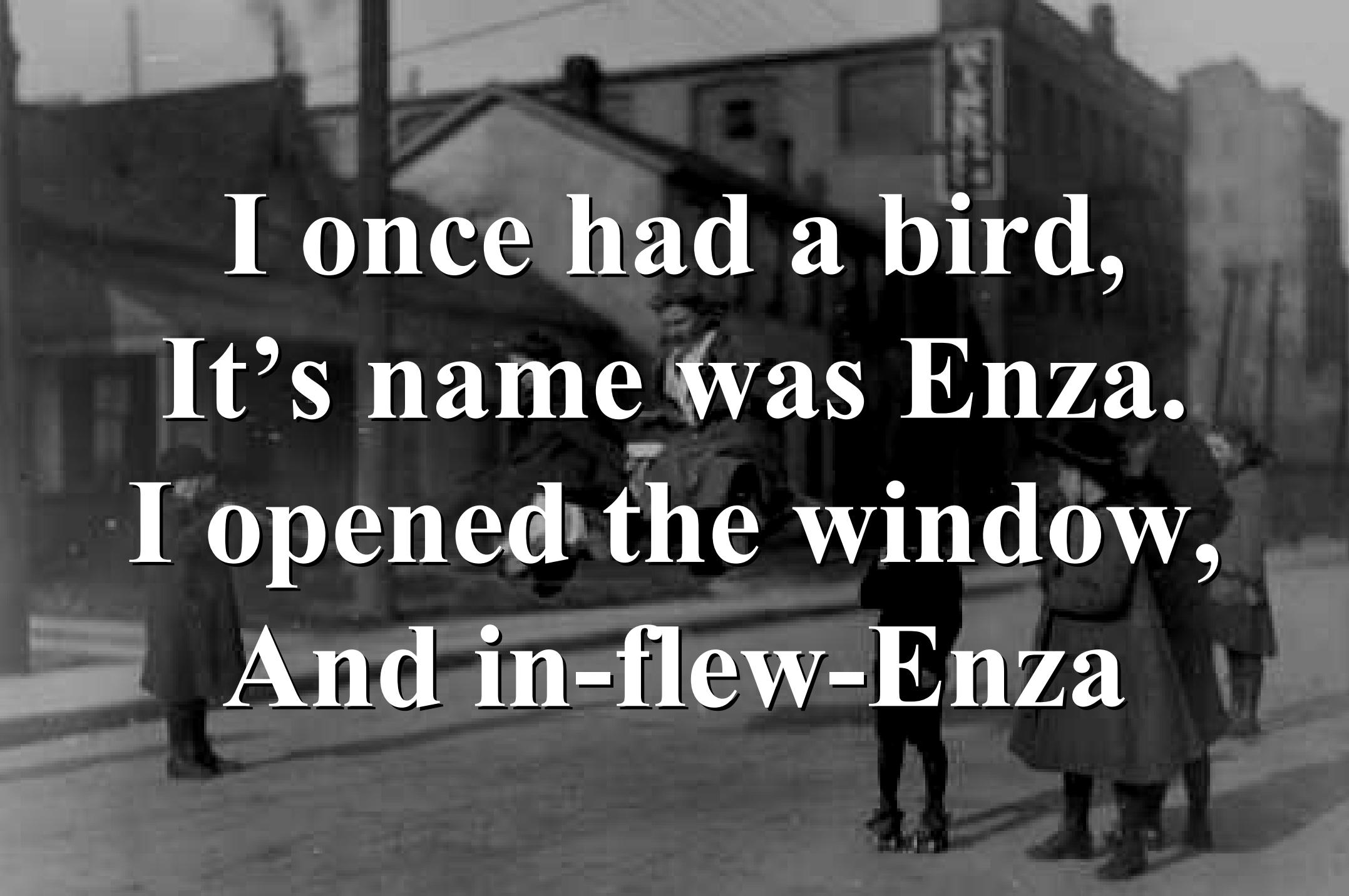




I once had a bird,
It's name was Enza.



I once had a bird,
It's name was Enza.
I opened the window,

A black and white photograph showing a person from behind, standing on a porch or balcony. They are looking out over a garden with several trees and shrubs. In the upper left corner of the image, there is a small, rectangular, framed object that looks like a birdhouse or a picture frame hanging on a wall.

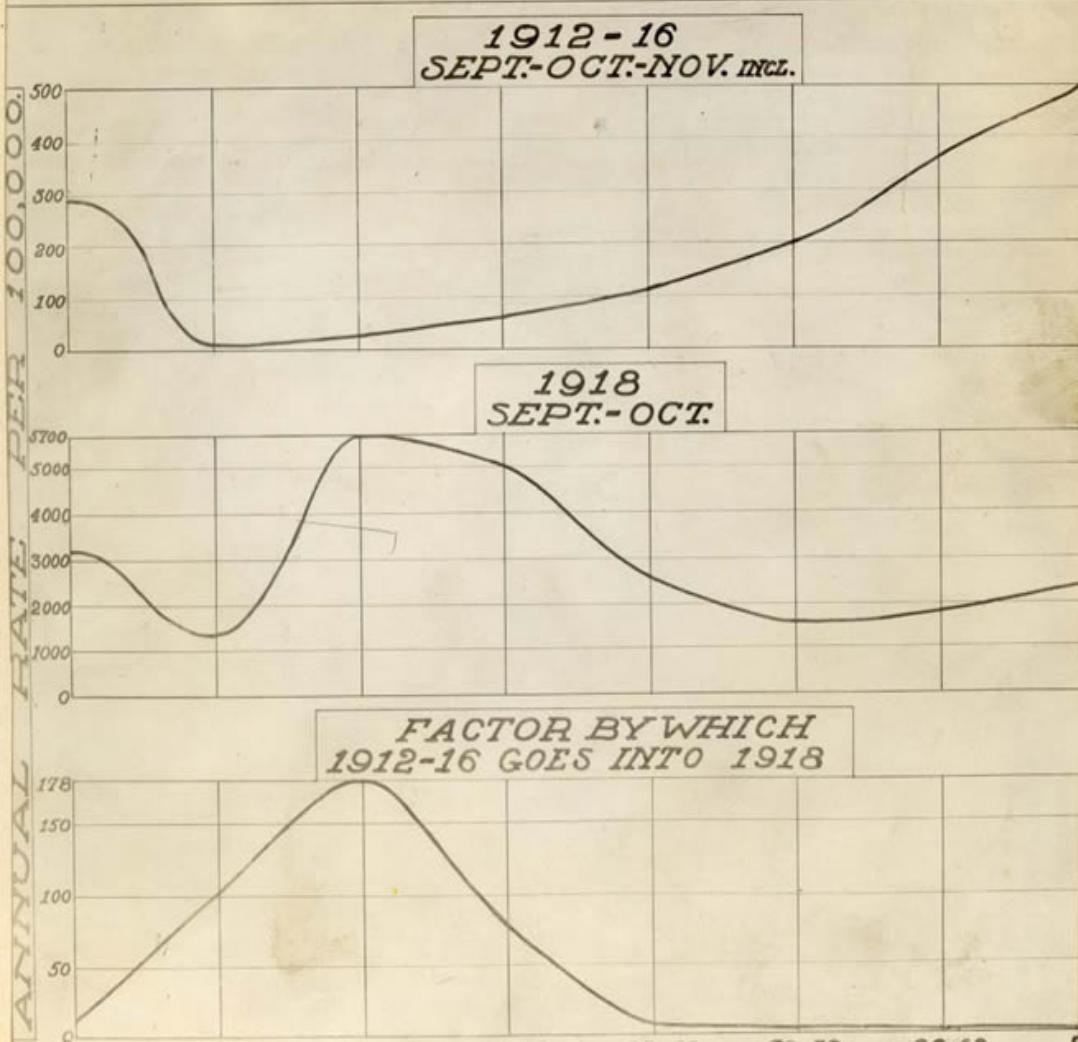
I once had a bird,
It's name was Enza.

I opened the window,
And in-flew-Enza

EXCESS MORTALITY IN U.S. CITIES DURING INFLUENZA EPIDEMIC PERCENT OF POPULATION DYING

CITY	1918-1919						
	SEPT. 8 - NOV. 23 10 WEEKS		NOV. 24 - FEB. 1 10 WEEKS		FEB. 2 - MAR 29 8 WEEKS		TOTAL 28 WEEKS
	0	.2	.4	.6	.8		
PHILADELPHIA	.69					.01	.73
FALL RIVER	.59					.05	.68
PITTSBURGH	.59					.12	.77
BALTIMORE	.57					.03	.60
SYRACUSE	.55					.02	.58
NASHVILLE	.55					.16	.83
BOSTON	.50					.12	.62
NEW HAVEN	.49					.13	.61
NEW ORLEANS	.49					.21	.71
ALBANY	.48					.03	.53
BUFFALO	.47					.10	.61
WASHINGTON	.45					.18	.54
LOWELL	.44					.10	.56
SAN FRANCISCO	.42					.31	.74
CAMBRIDGE	.39					.12	.50
NEWARK	.38					.11	.53
PROVIDENCE	.38					.13	.53
RICHMOND	.35					.18	.55
DAYTON	.33					.02	.55
OAKLAND	.33					.22	.37
CHICAGO	.32					.09	.56
NEW YORK	.30					.09	.46
CLEVELAND	.27					.11	.47
LOS ANGELES	.27					.26	.42
MEMPHIS	.25					.02	.55
ROCHESTER	.25					.12	.37
KANSAS CITY	.25					.27	.40
DENVER	.24					.32	.63
CINCINNATI	.22					.13	.46
OMAHA	.22					.20	.43
LOUISVILLE	.19					.04	.37
ST. PAUL	.19					.13	.34
COLUMBUS	.19					.15	.41
PORTLAND	.18					.22	.42
TOLEDO	.17					.08	.17
MINNEAPOLIS	.17					.11	.24
SEATTLE	.16					.18	.36
INDIANAPOLIS	.15					.09	.31
BIRMINGHAM	.15					.15	.29
MILWAUKEE	.15					.18	.37
ST. LOUIS	.12					.18	.34
SPOKANE	.11					.13	.25
ATLANTA	.07					.13	.19
GRAND RAPIDS	.04					.12	.19

AGE DISTRIBUTION OF DEATHS FROM INFLUENZA AND PNEUMONIA AT BOSTON



EXCESS MORTALITY IN U.S. CITIES DURING INFLUENZA EPIDEMIC PERCENT OF POPULATION DYING

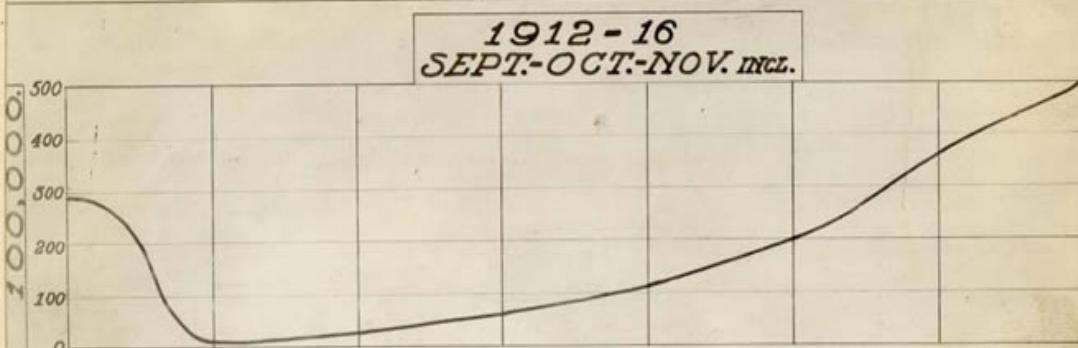
CITY	1918-1919				
	SEPT. 8 - NOV. 23 10 WEEKS	NOV. 24 - FEB. 1 10 WEEKS	FEB. 2 - MAR 29 8 WEEKS	TOTAL 28 WEEKS	
	0	.2	.4	.6	.8
PHILADELPHIA	.69				
FALL RIVER	.59				
PITTSBURGH	.59				
BALTIMORE	.57				
SYRACUSE	.55				
NASHVILLE	.55				
BOSTON	.50				
NEW H					
NEW C					
ALBAN					
BUFFALO					
WASHIN					
LOWELL					
SAN FR					
CAMBR					
NEWAI					
PROVII					
RICHM					
DAYTO					
OAKLA					
CHICAG					
NEW Y					
CLEVE					
LOS AN					
MEMPHIS	.25				
ROCHESTER	.25				
KANSAS CITY	.25				
DENVER	.24				
CINCINNATI	.22				
OMAHA	.22				
LOUISVILLE	.19				
ST. PAUL	.19				
COLUMBUS	.19				
PORTLAND	.18				
TOLEDO	.17				
MINNEAPOLIS	.17				
SEATTLE	.16				
INDIANAPOLIS	.15				
BIRMINGHAM	.15				
MILWAUKEE	.15				
ST. LOUIS	.12				
SPOKANE	.11				
ATLANTA	.07				
GRAND RAPIDS	.04				

SEPT. 8 - NOV. 23
10 WEEKS

0 .2 .4 .6 .8

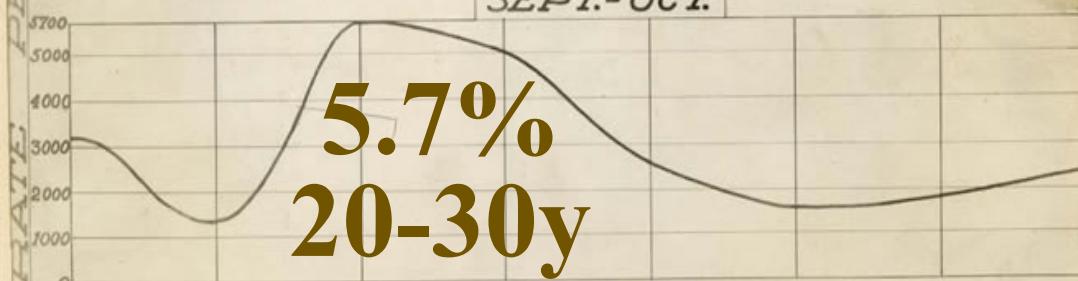
.69

AGE DISTRIBUTION OF DEATHS FROM INFLUENZA AND PNEUMONIA AT BOSTON



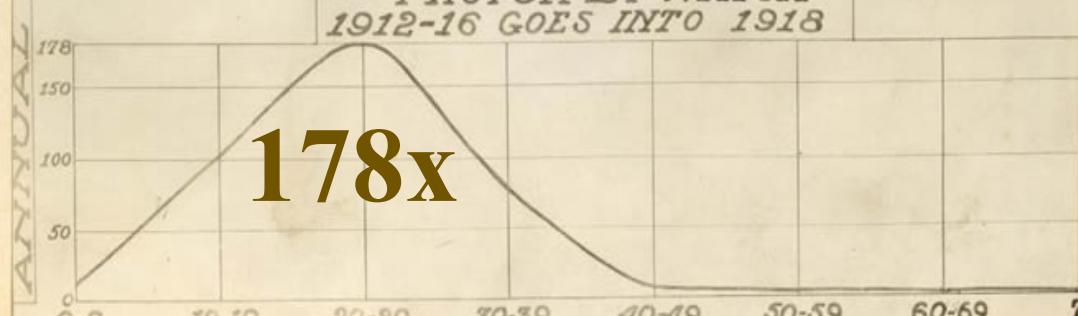
1918
SEPT.-OCT.

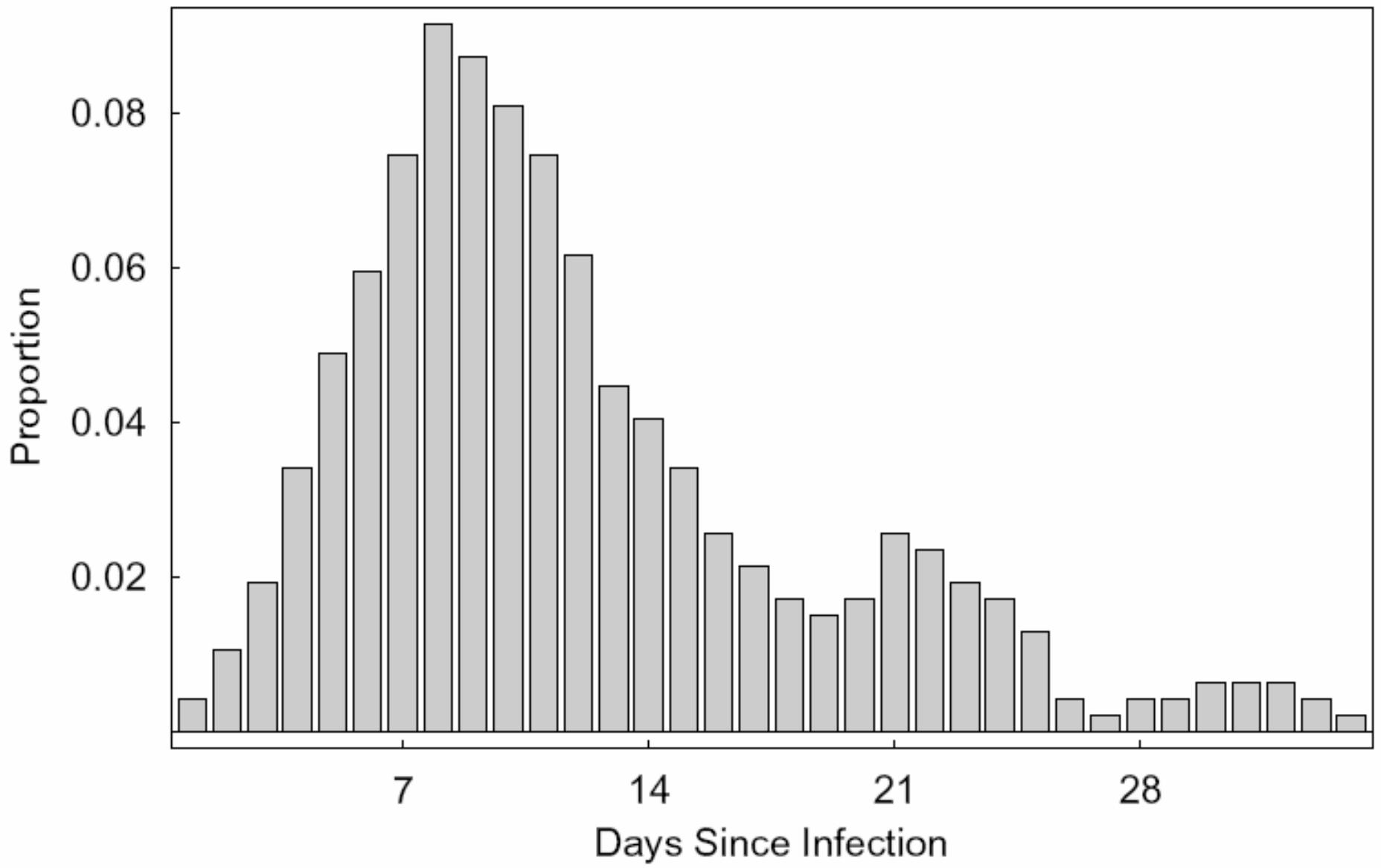
5.7%
20-30y



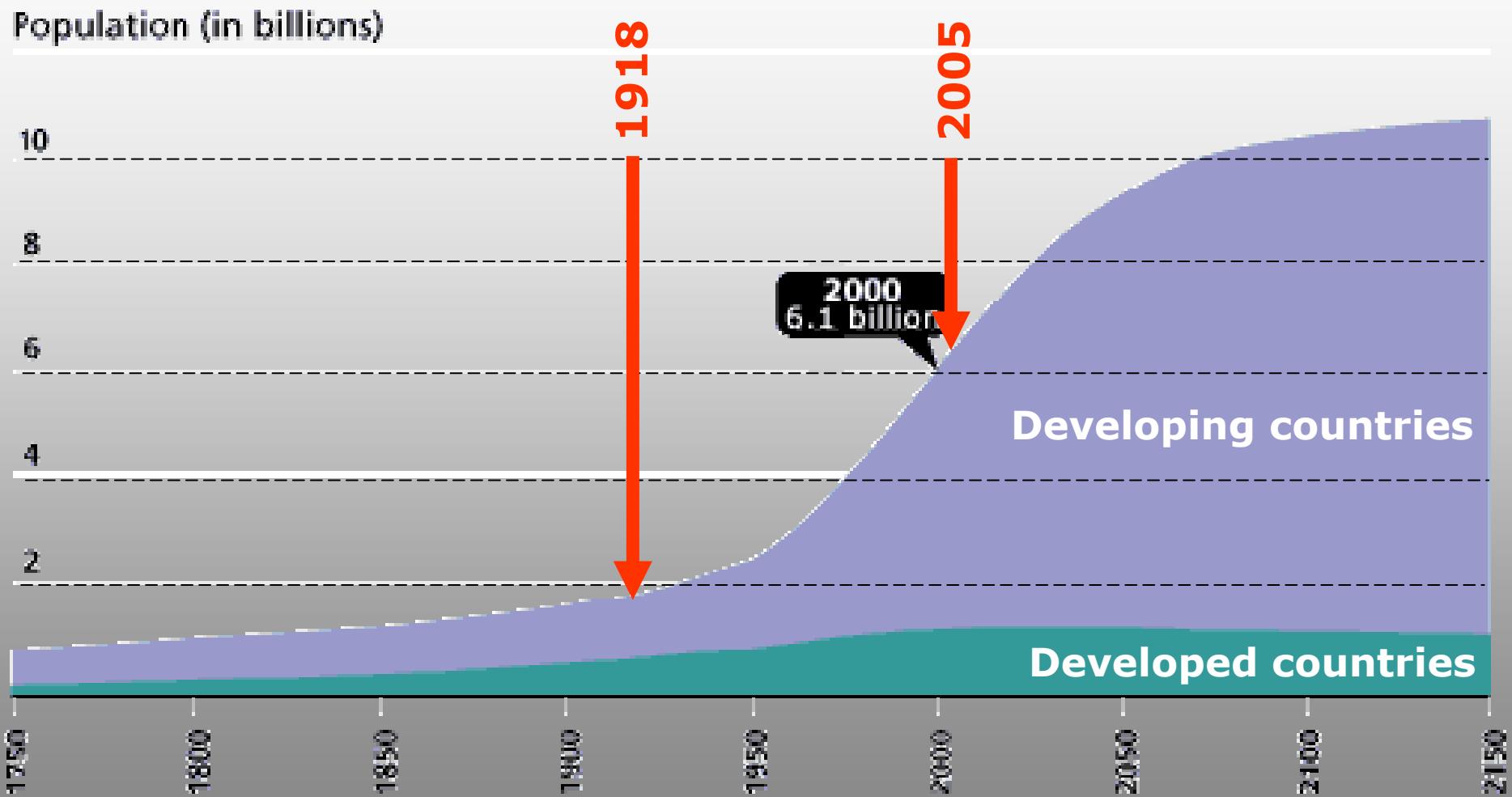
FACTOR BY WHICH
1912-16 GOES INTO 1918

178x

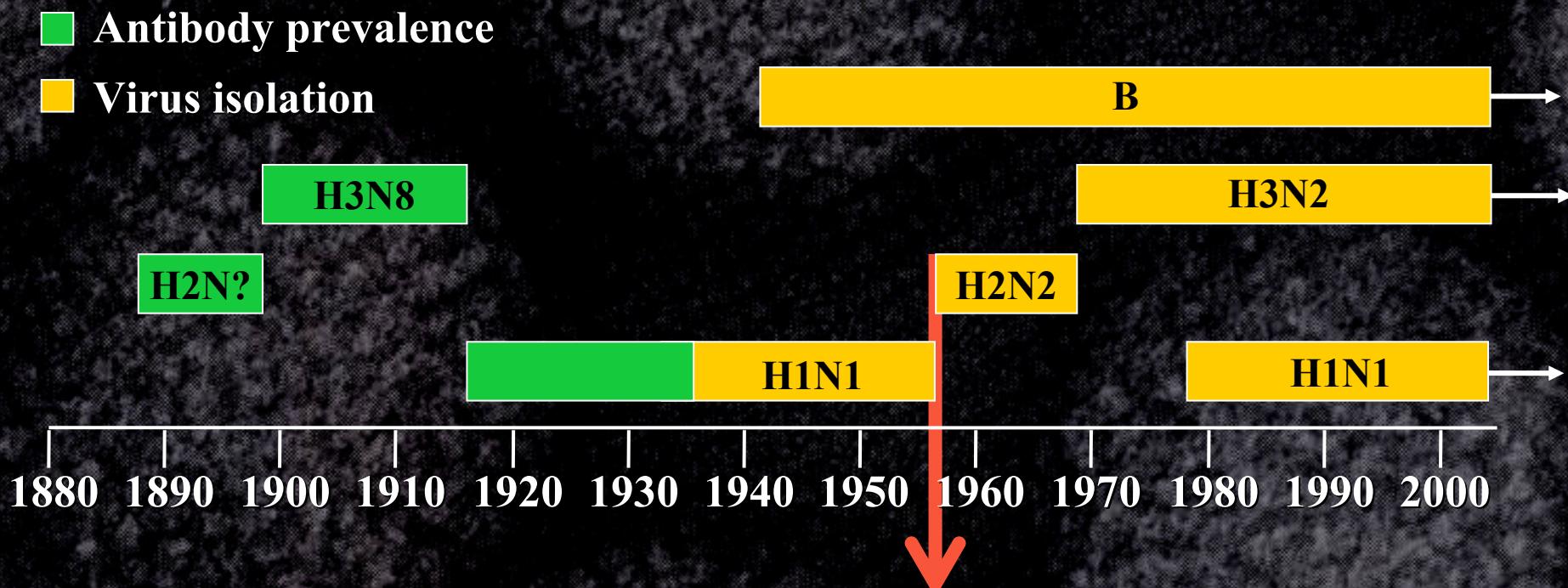




POPULATION GROWTH 1750-2150

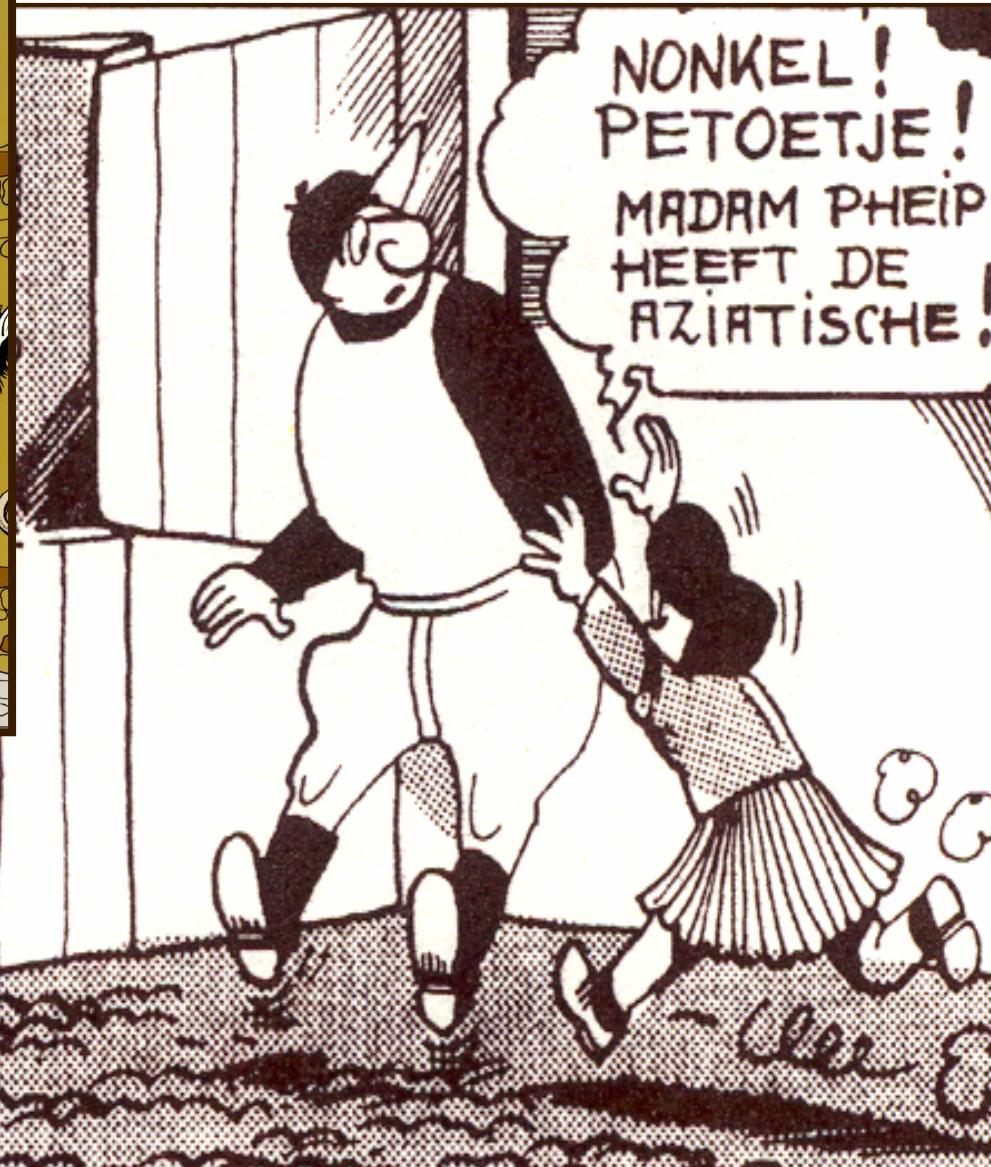


INFLUENZA VIRUSES



1957 H2N2:
Asian flu

HET GEHEIM VAN BAKKENDOEN



JAH, DE
AZIATISCHE
GRIEP
IS OP ONZE
HELDEN
NEERGESTORT.

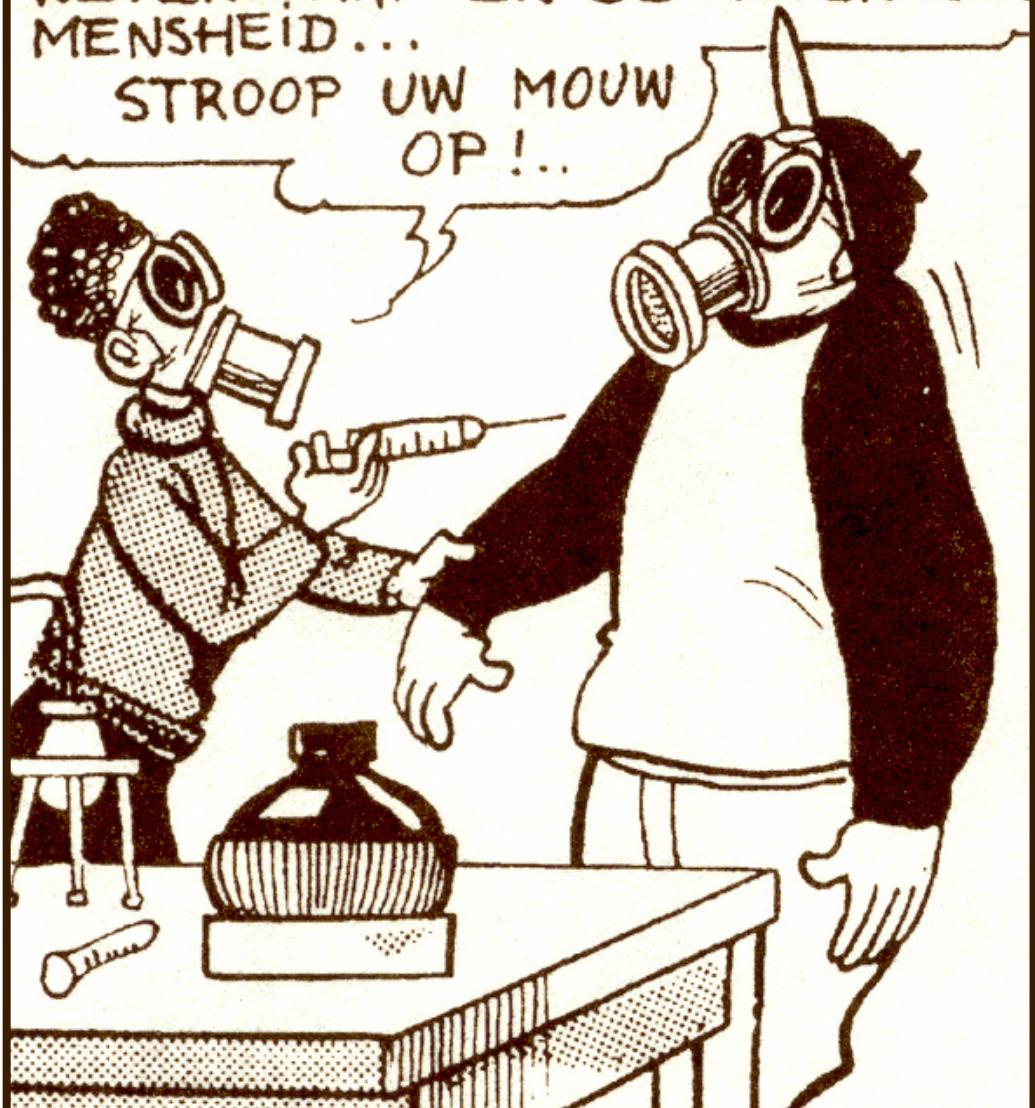
MENEER PHEIP
KWAM ER MEE
NAAR HUIS
VAN ZIJN
STAMCAFE.
DAARNA KREEG
ZE MADAM
PHEIP TE
PAKKEN
EN NU...

PROBEER EEN ANTI- AZIATIEKER.
ISOLEER DE VIRUS. DOE IETS,
GE ZIJT GELEERD GENOEG.
IK ZAL CONTROLEREN!

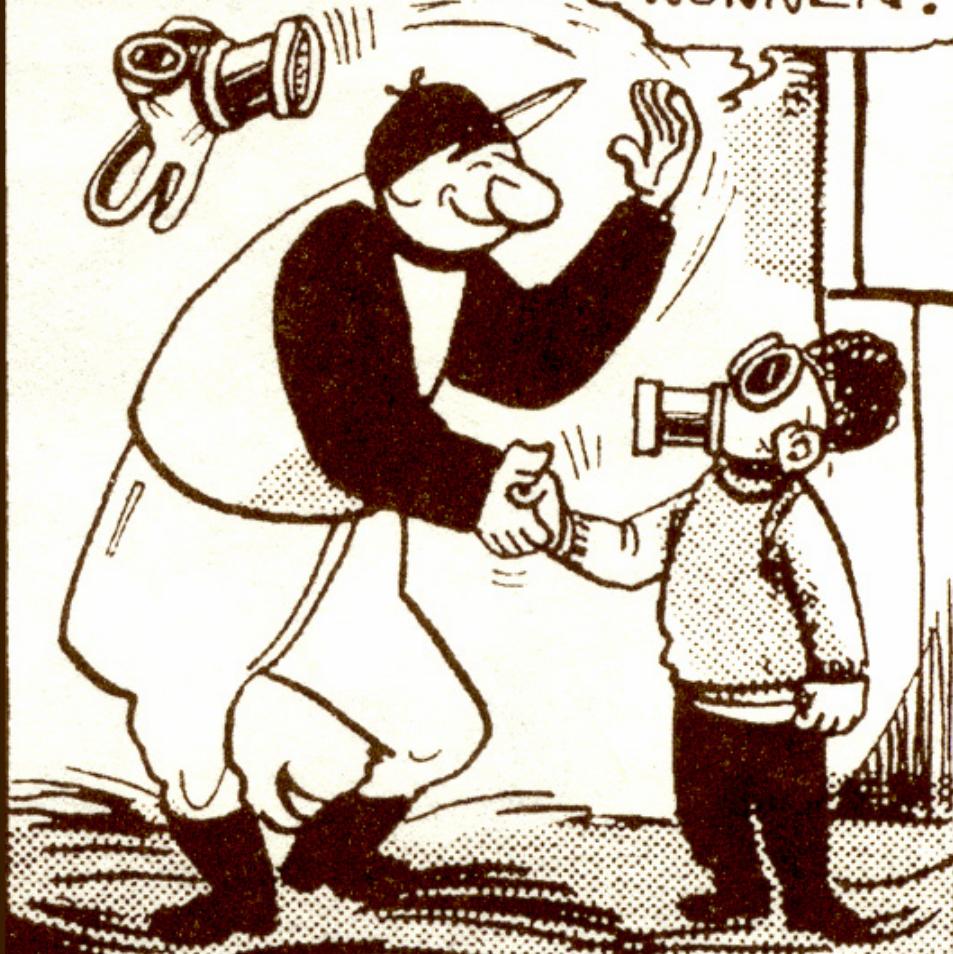


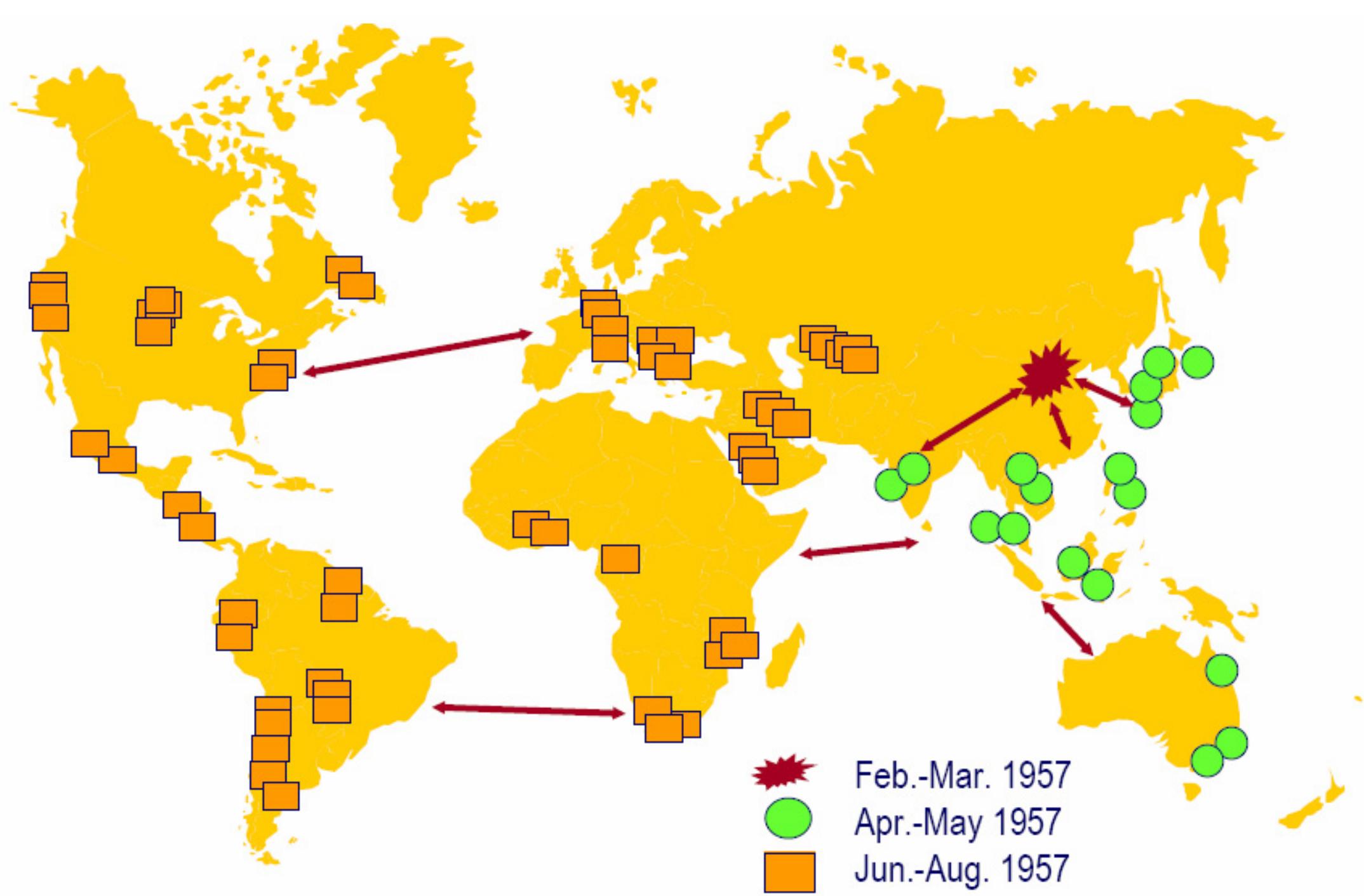
NONKEL, IN HET BELANG VAN DE
WETENSCHAP EN DE GETEISTERDE
MENSHEID...

STROOP UW MOUW
OP!..

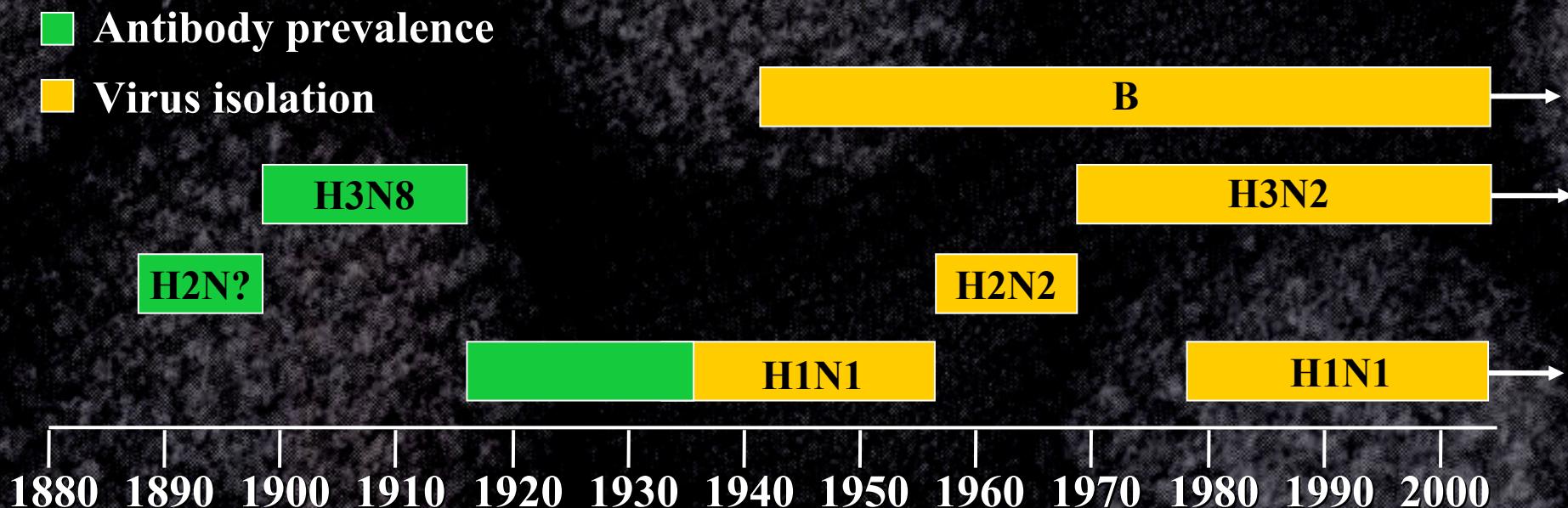


PROFICIAT PETOETJE. GE WORDT,
NOG EEN GROOT GELEERDE.
DE MASKERS MOGEN WE WEG-
WERPEN, WE HEBBEN DE
AZIATISCHE GRIEP OVER-
WONNEN!





INFLUENZA VIRUSES



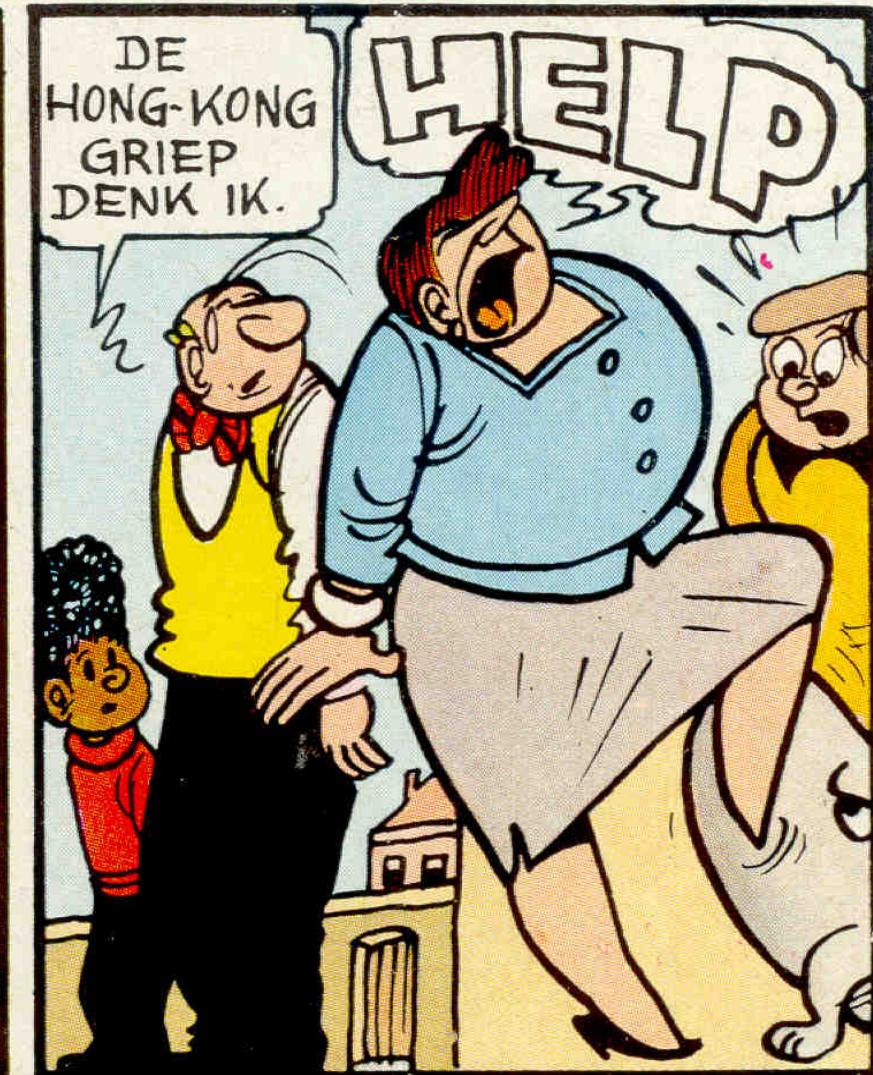
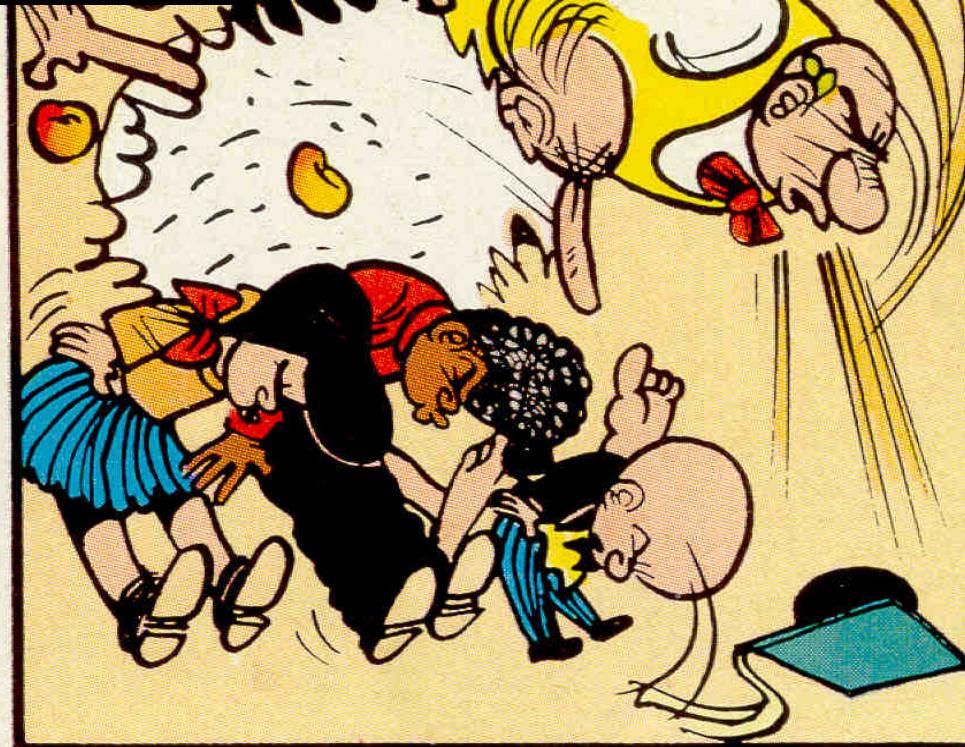
1968 H3N2:
Hong Kong flu



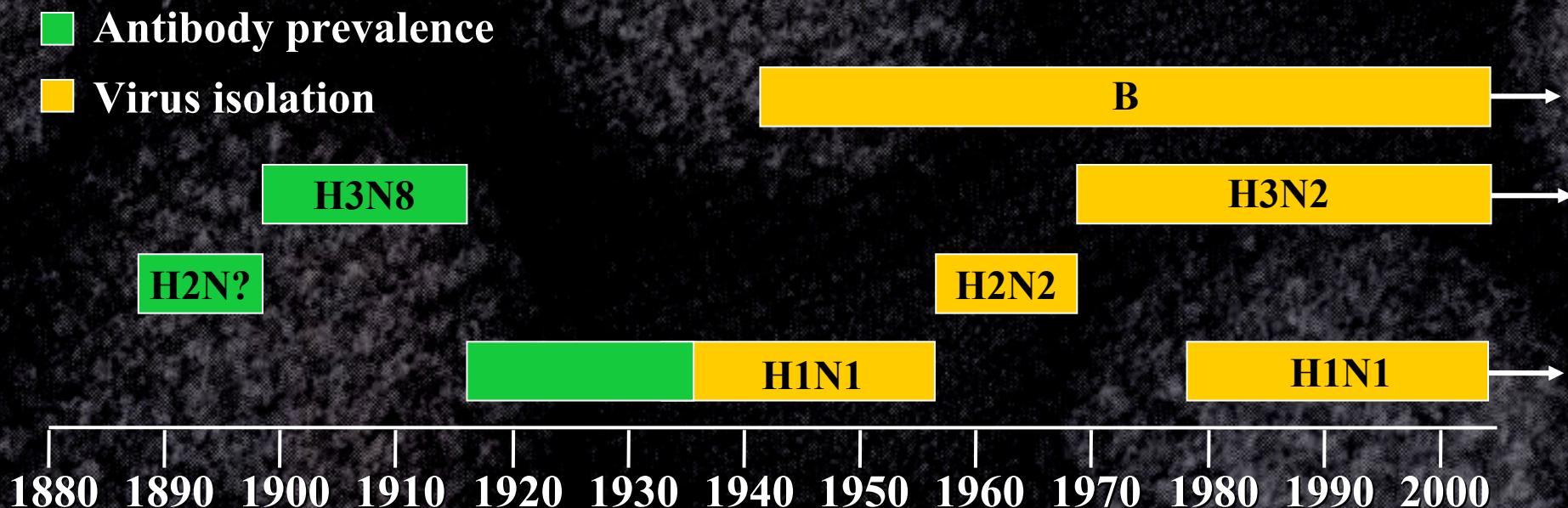
Marc Sleen

DE AVONTUREN VAN NERO
EN CO.

TOFFE THEO



INFLUENZA VIRUSES



1977 H1N1:
Russian flu

EDICAL ALERT!

AL JIMSON / Westing World News

A new killer flu is poised to sweep America — and experts fear it to claim millions of victims from all walks of life!

The enormous fatality rate will come because of increases in world population, a larger — and poorer — world population, and the fact that most no longer possess antibodies to deadly flu like the one that killed 20 million people around the world in 1918, scientists say.

"It's a major threat," says Dominick Faziozzi of the National Institute of Allergy and Infectious Diseases. "It would wreak tremendous damage on the U.S. and around the world."

Such a giant flu — called a pandemic — has historically only periodically attacked mankind in killer waves. Last major flu pandemic was the Hong Kong flu of 1968, which killed 70,000 people in the U.S. alone. The next one, experts fear, could be far, far worse.

"A new subtype of virus has started to run through the population," warning Lorraine Johnson, "and it would cause severe disease and widespread infections. No one would have any immunity to it."

Health authorities are especially worried because pandemics like the 1918 one sweep not only the young and the very-old, but also healthy people in the prime of life. Experts are worried that many victims remain hours of infection.

Witnesses to the deadly 1918 influenza recall a healthy young man who was a streetcar in Chicago and died when the trolley had gone six blocks. A tall of a woman who boarded a train in New York in seemingly poor health — and landed over dead in a half hour.

Though attempts have been made to stop newly emerging flu viruses from infecting populations in Africa, experts say it's inevitable that an unexpected and hitherto unknown virus will eventually develop and spread. When it does, there will be helpless victims despite all the international breakthroughs.

"I think we could do something about it if the 1918 virus came back again," says virologist William D. Bellamy of New York Medical College, "but of a breakthrough in the viruses themselves. We never vaccinate the population all

KILLER STRIKE



TENT CITIES were built when hospitals couldn't handle the millions who were stricken during the 1918 pandemic.

SUPERBUG INNOCENT

ININCREDIBLE PHOTO shows an influenza virus magnified 132,000 times. Experts believe a deadly new mutation could emerge any day, guaranteeing the worst epidemic since.



FLU ABOUT AMERICA!

WILL CLAIM THE LIVES OF MILLIONS OF VICTIMS, WARN WORRIED EXPERTS

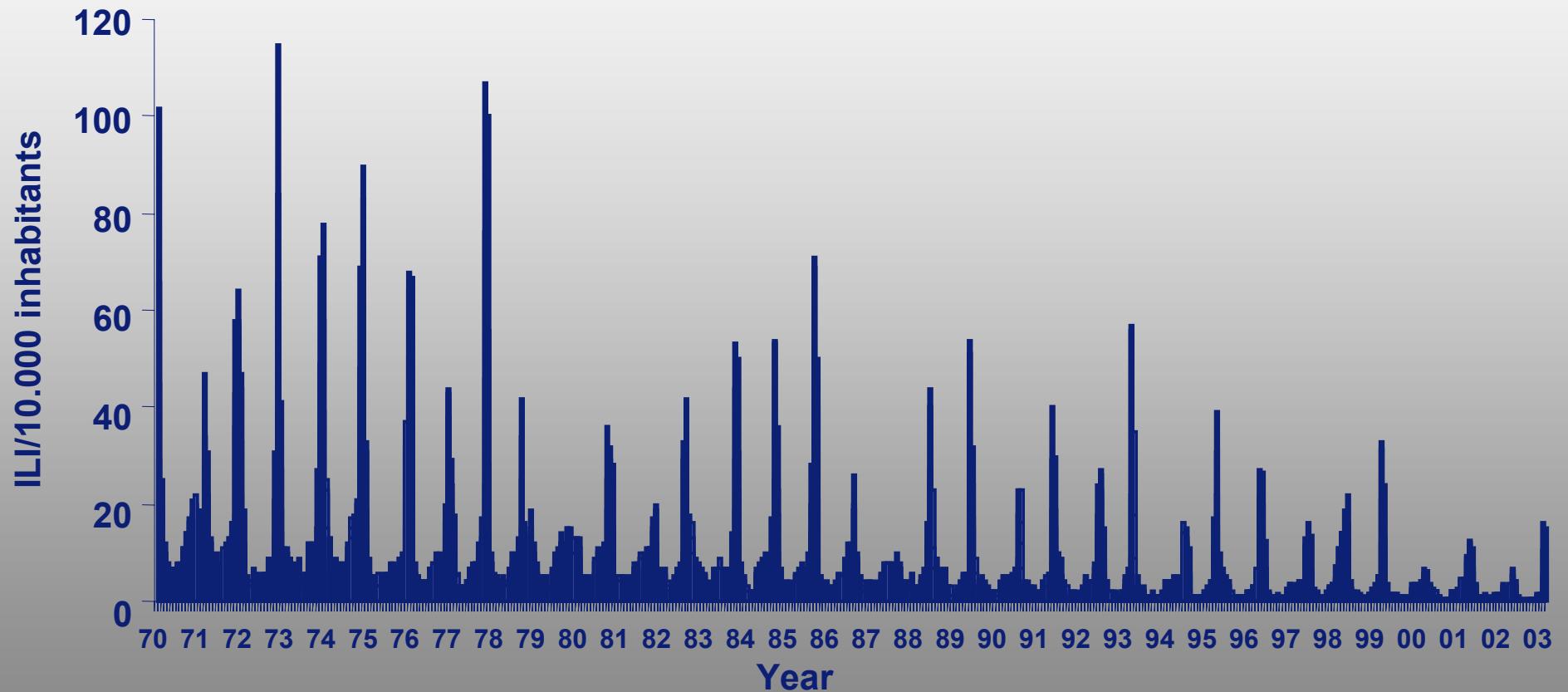


HELPLESS DOCTORS were unable to save the more than 20 million people who died during the 1918 outbreak.

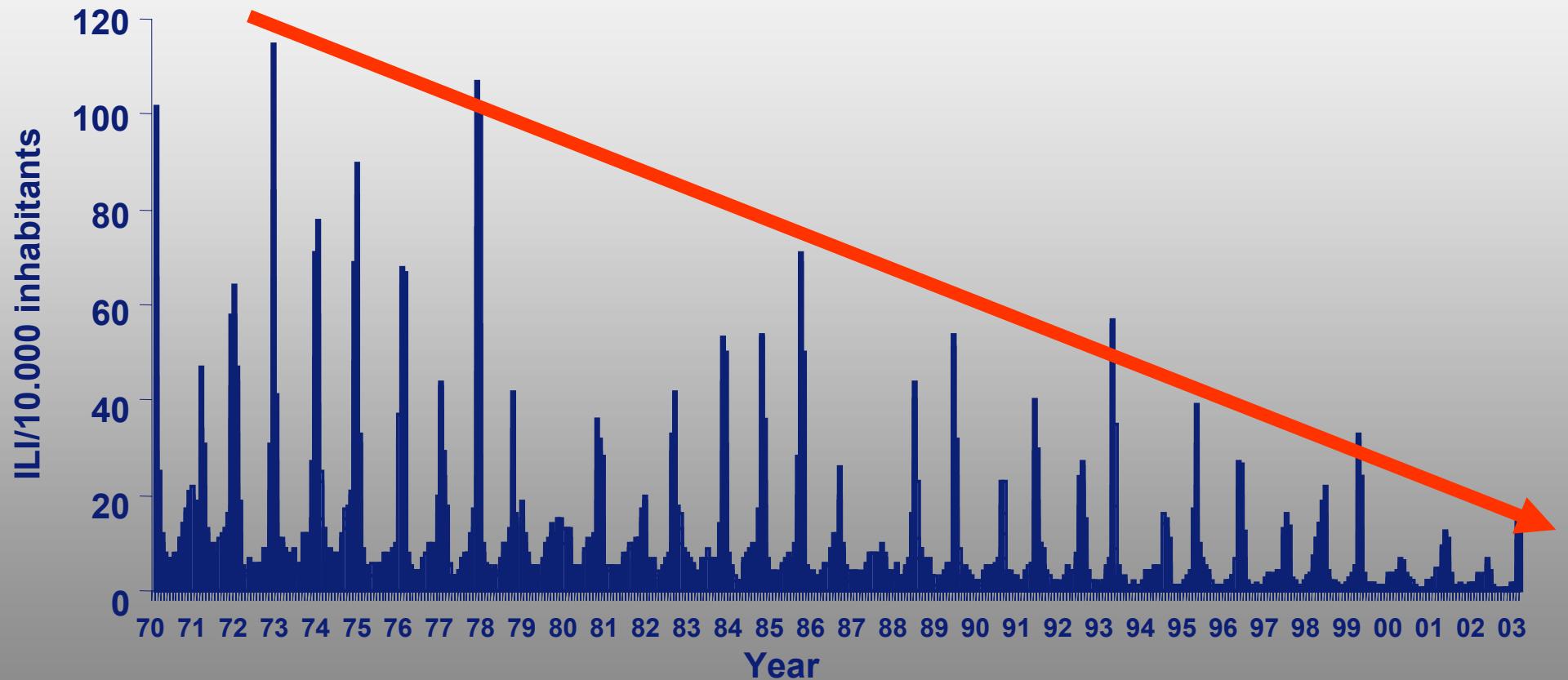


HOSPITALS ACROSS THE COULD

Influenza-like illness in the Netherlands per week per 10.000 inhabitantis



Influenza-like illness in the Netherlands per week per 10.000 inhabitants

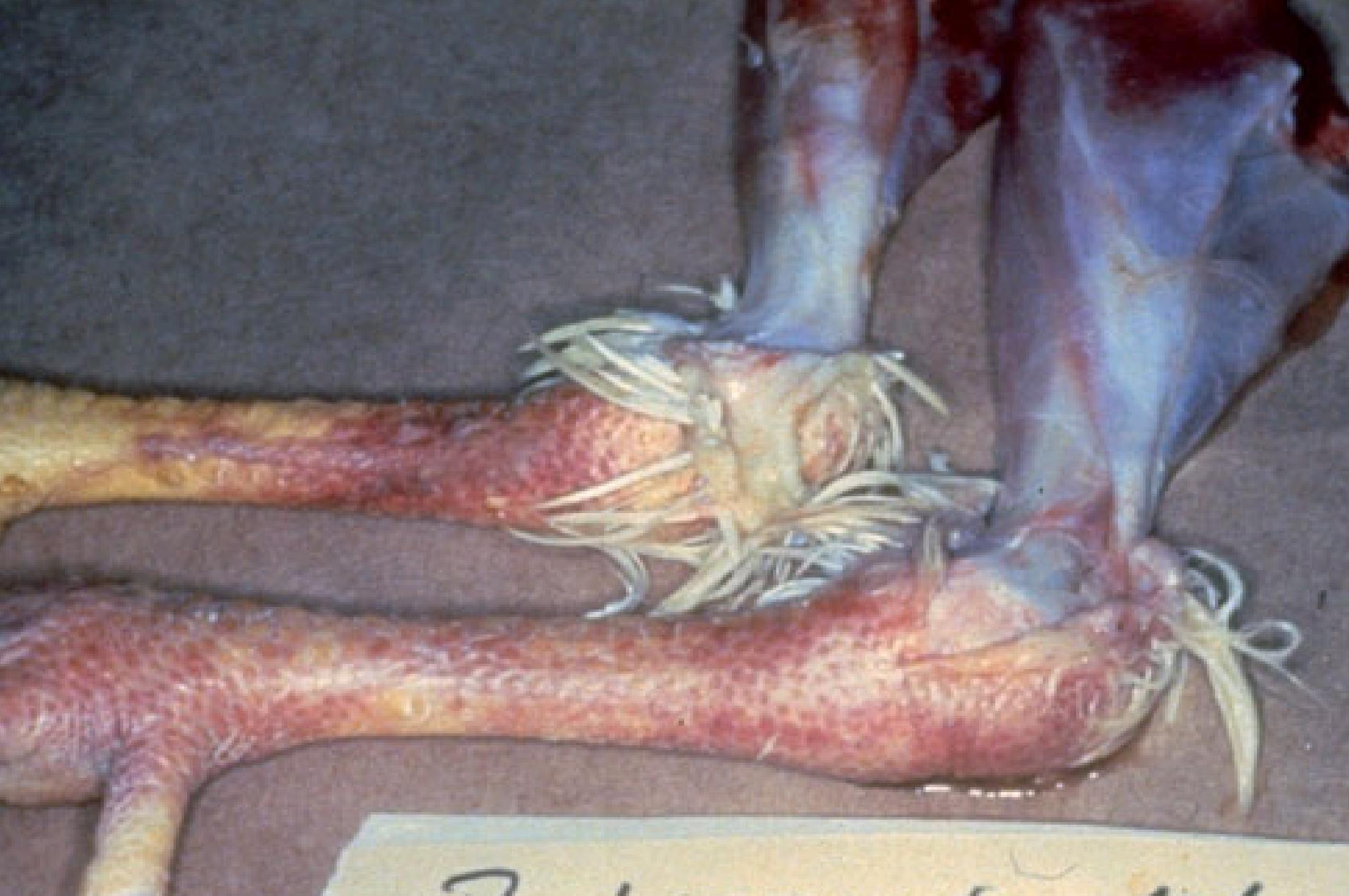












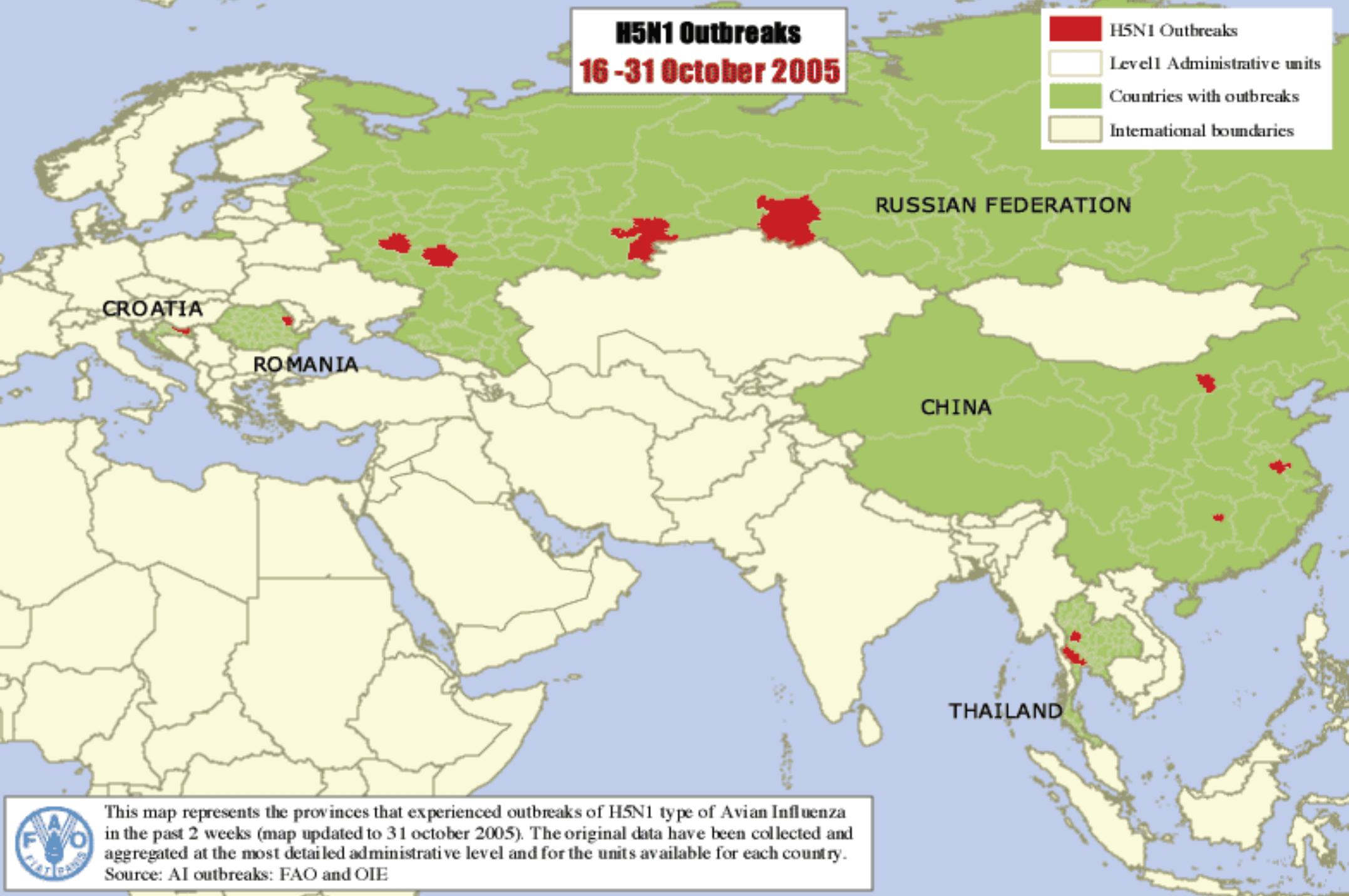
31





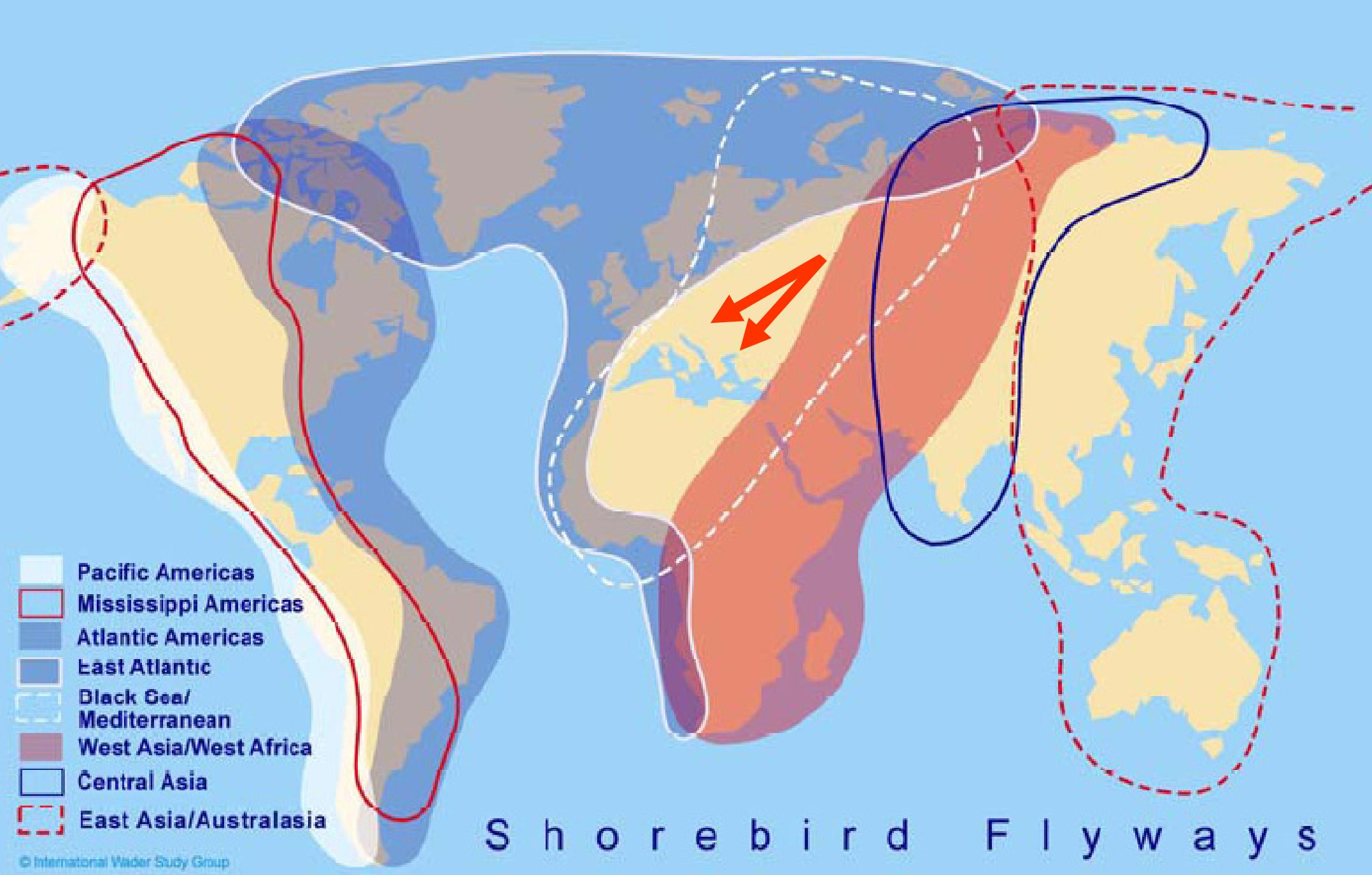
H5N1 Outbreaks 16 -31 October 2005

- H5N1 Outbreaks
- Level 1 Administrative units
- Countries with outbreaks
- International boundaries



This map represents the provinces that experienced outbreaks of H5N1 type of Avian Influenza in the past 2 weeks (map updated to 31 October 2005). The original data have been collected and aggregated at the most detailed administrative level and for the units available for each country.
Source: AI outbreaks: FAO and OIE

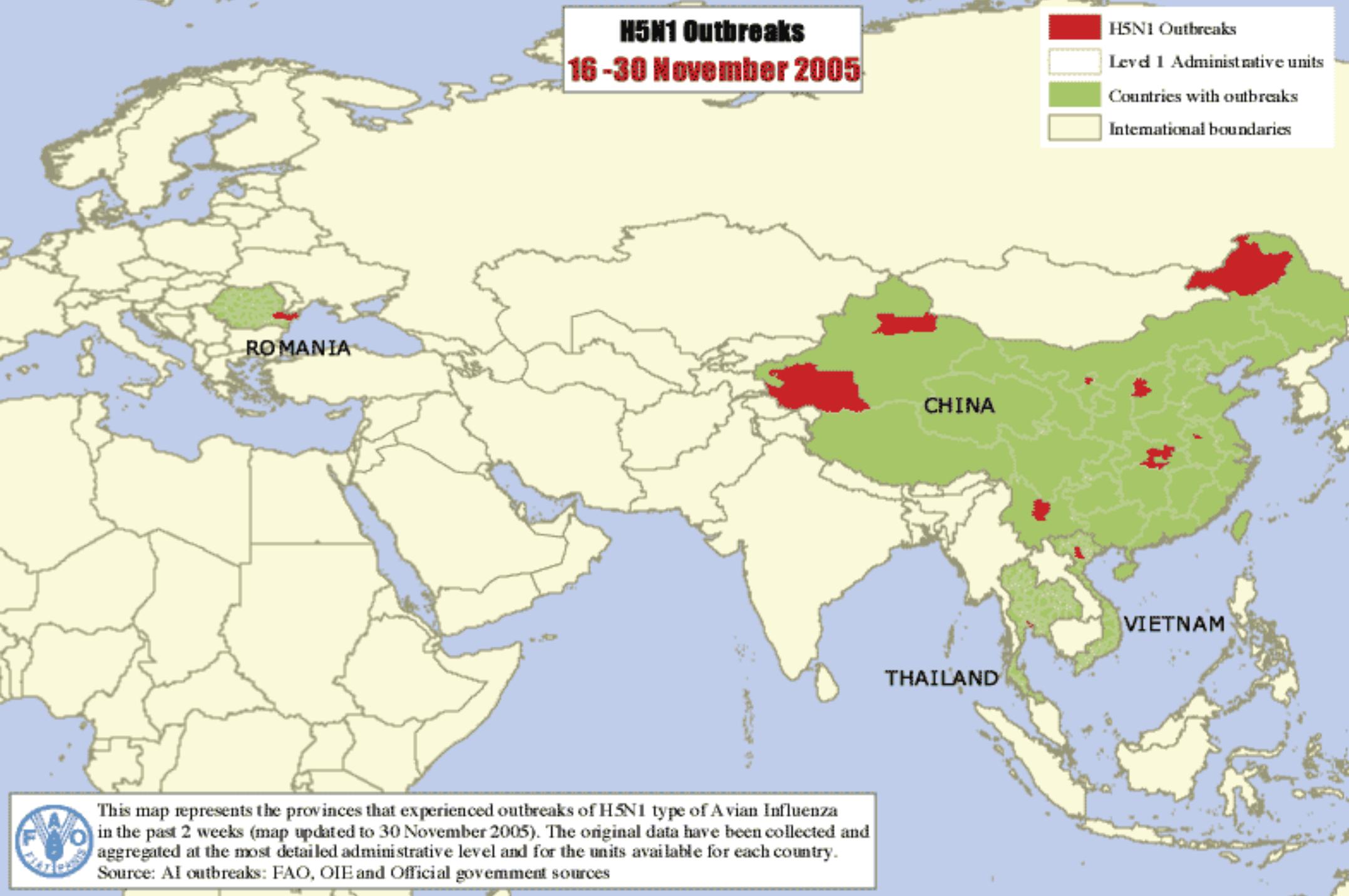




H5N1 Outbreaks

16 - 30 November 2005

- H5N1 Outbreaks
- Level 1 Administrative units
- Countries with outbreaks
- International boundaries



This map represents the provinces that experienced outbreaks of H5N1 type of Avian Influenza in the past 2 weeks (map updated to 30 November 2005). The original data have been collected and aggregated at the most detailed administrative level and for the units available for each country.
Source: AI outbreaks: FAO, OIE and Official government sources



Cumulative Number of Confirmed Human Cases of Avian Influenza A/
(H5N1) Reported to WHO

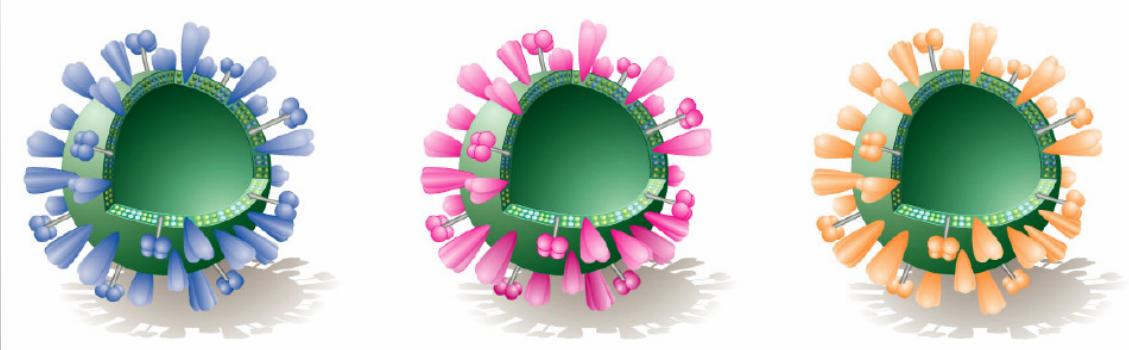
6 December 2005

Date of onset	Cambodia		China		Indonesia		Thailand		Viet Nam		Total	
	cases	deaths	cases	deaths	cases	deaths	cases	deaths	cases	deaths	cases	deaths
26.12.03- 10.03.04	0	0	0	0	0	0	12	8	23	16	35	24
19.07.04- 08.10.04	0	0	0	0	0	0	5	4	4	4	9	8
16.12.04- to date	4	4	3	2	13	8	4	1	66	22	90	37
Total	4	4	3	2	13	8	21	13	93	42	134	69

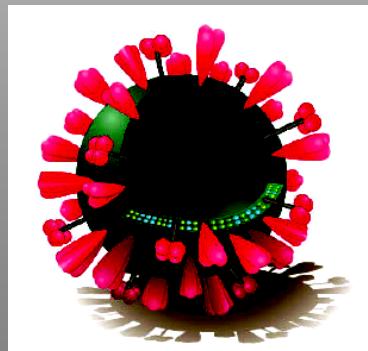
Total number of cases includes number of deaths.
WHO reports only laboratory-confirmed cases.



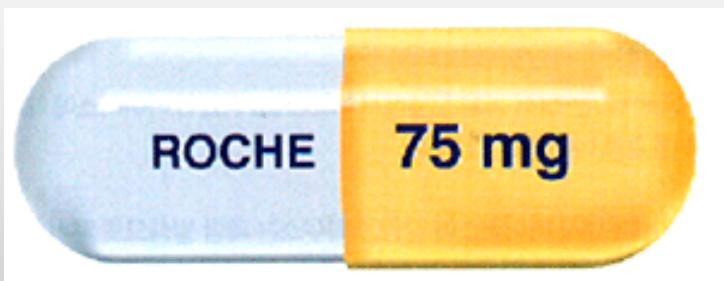
INCREASE IN VACCINATION COVERAGE TO 33% IN 2010



33%



100%



57.000 Euro



OKAY MR. BUSH...
WE ARE LISTENING.

IRAQ

SYRIA

N.KOREA

IRAN

CUBA

AXIS OF EVIL

SMALLPOX

AXIS  **OF** **EVIL**

SMALLPOX

MEASLES

AXIS  **OF** **EVIL**

700.000

SMALLPOX

MEASLES

MALARIA

2.000.000

700.000



AXIS OF EVIL

SMALLPOX

700.000

MEASLES

MALARIA

TBC

2.500.000

2.000.000



AXIS OF EVIL

SMALLPOX

700.000

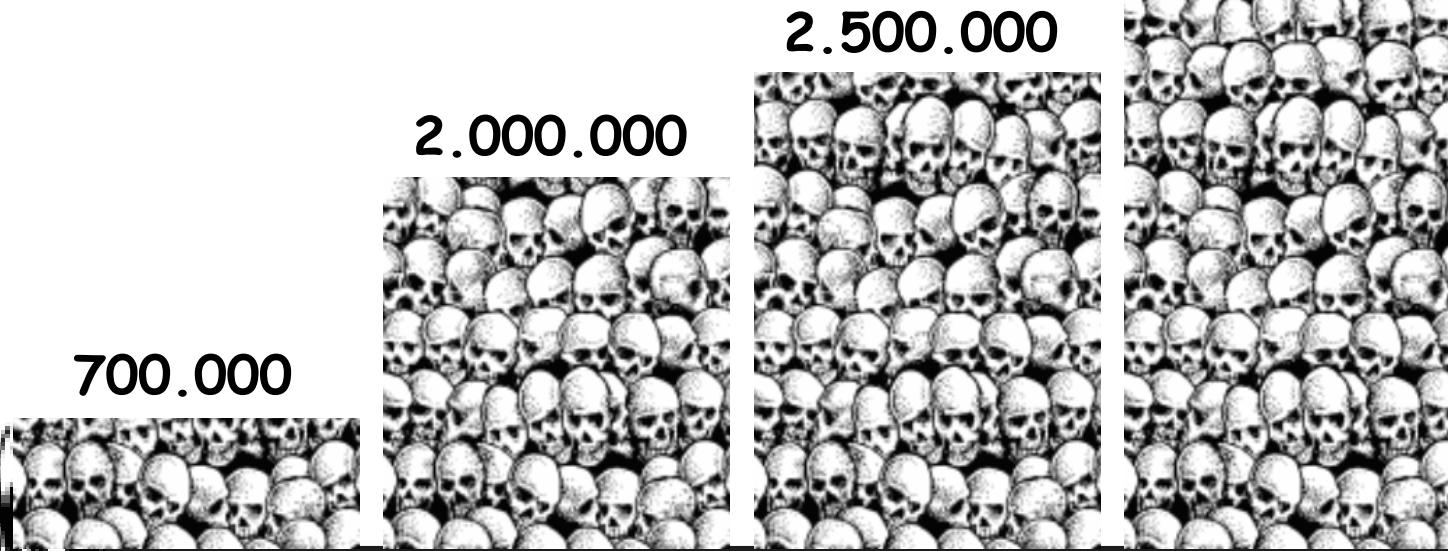
MEASLES

MALARIA

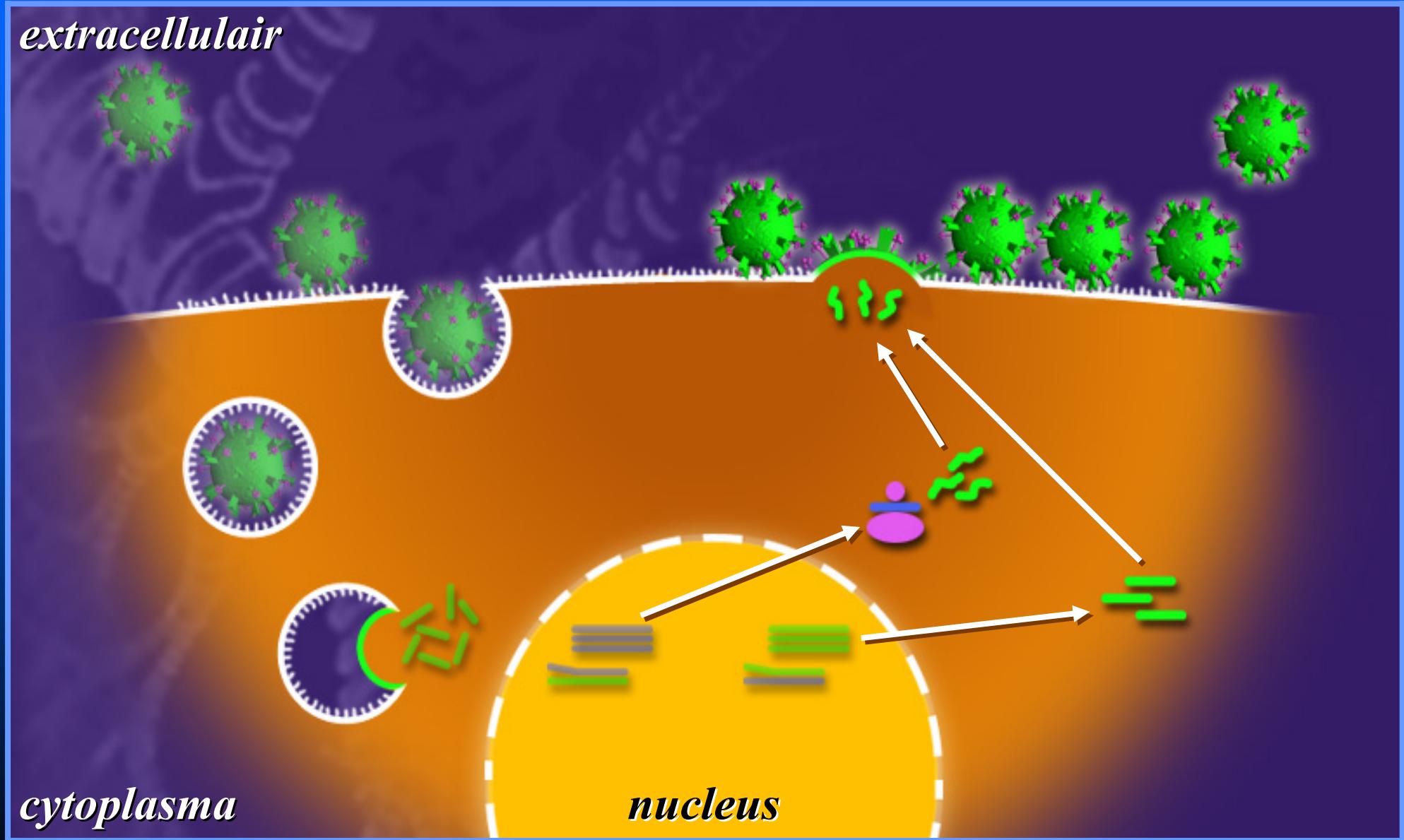
TBC

HIV

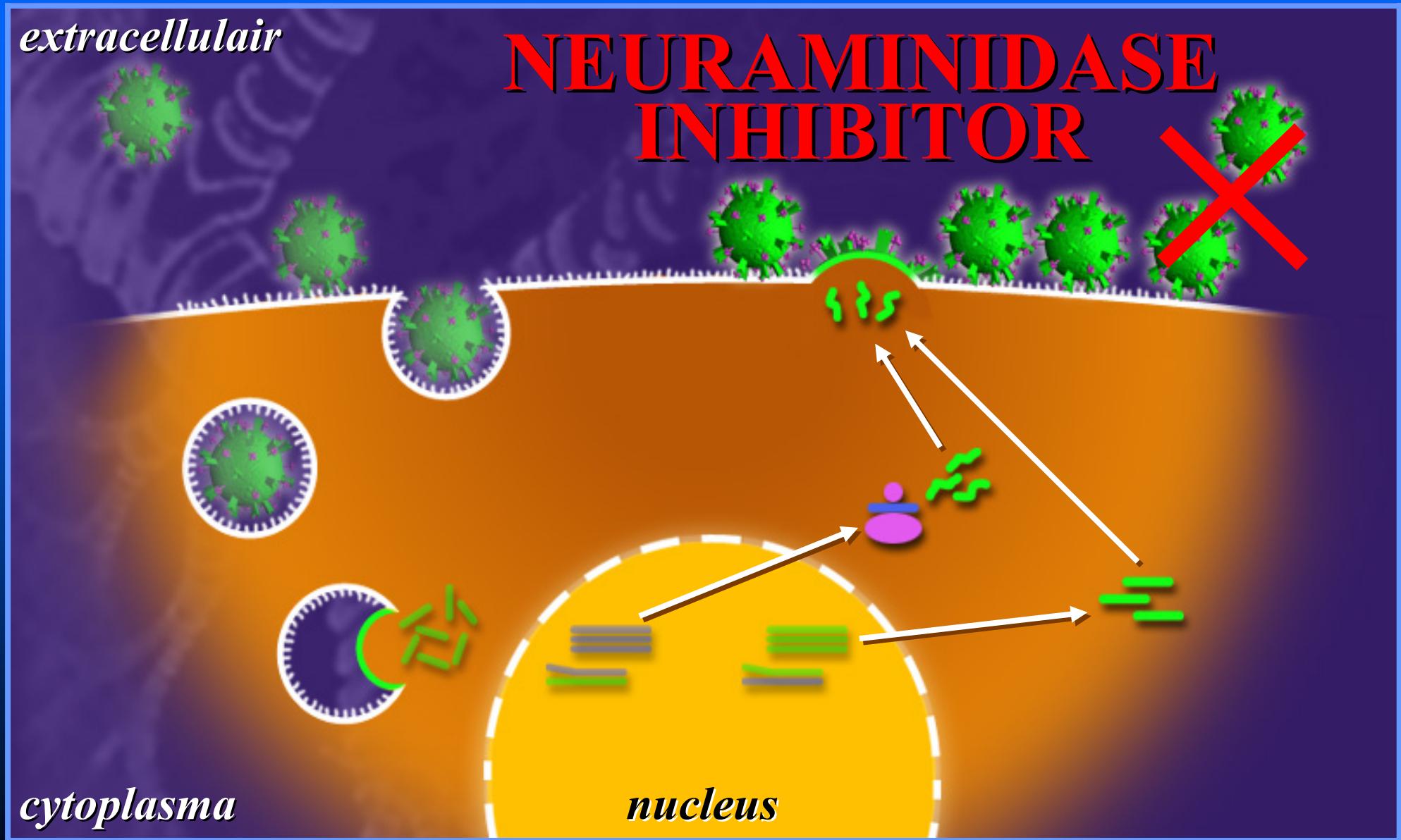
AXIS OF EVIL



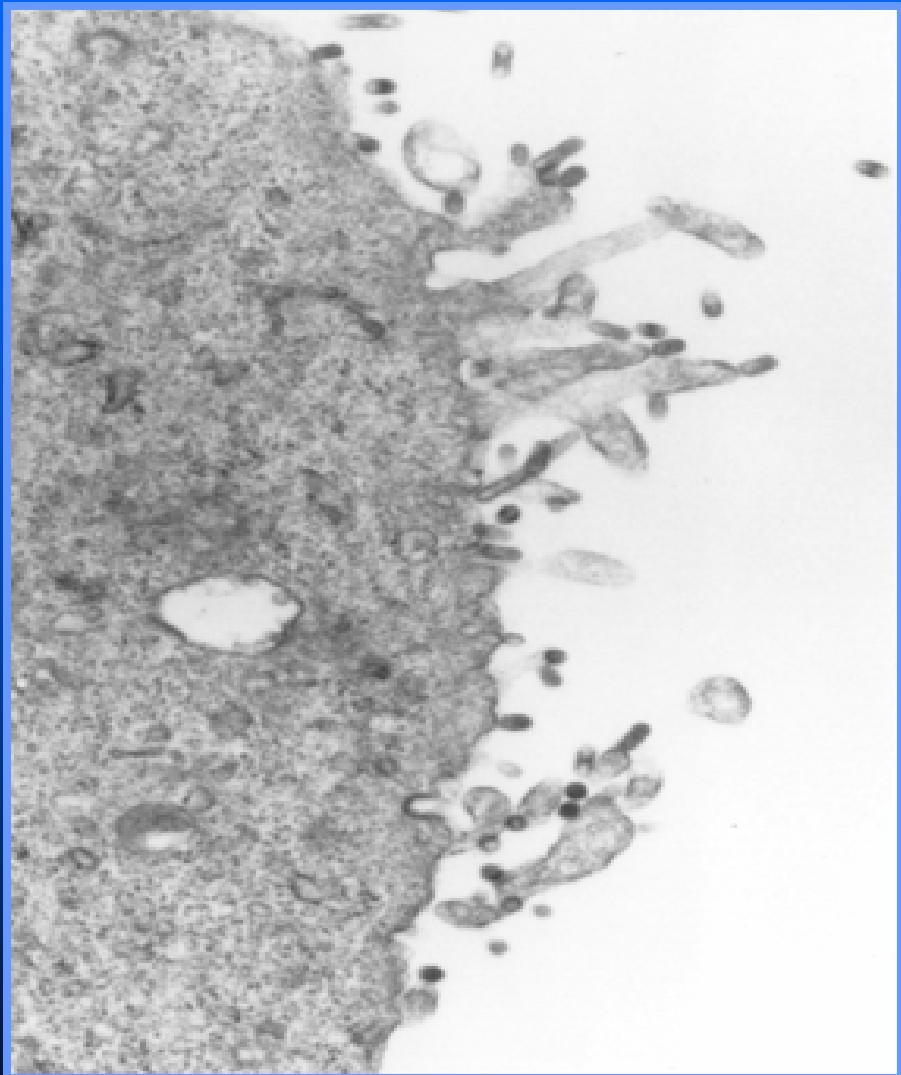
INFLUENZAVIRUS REPLICATIE



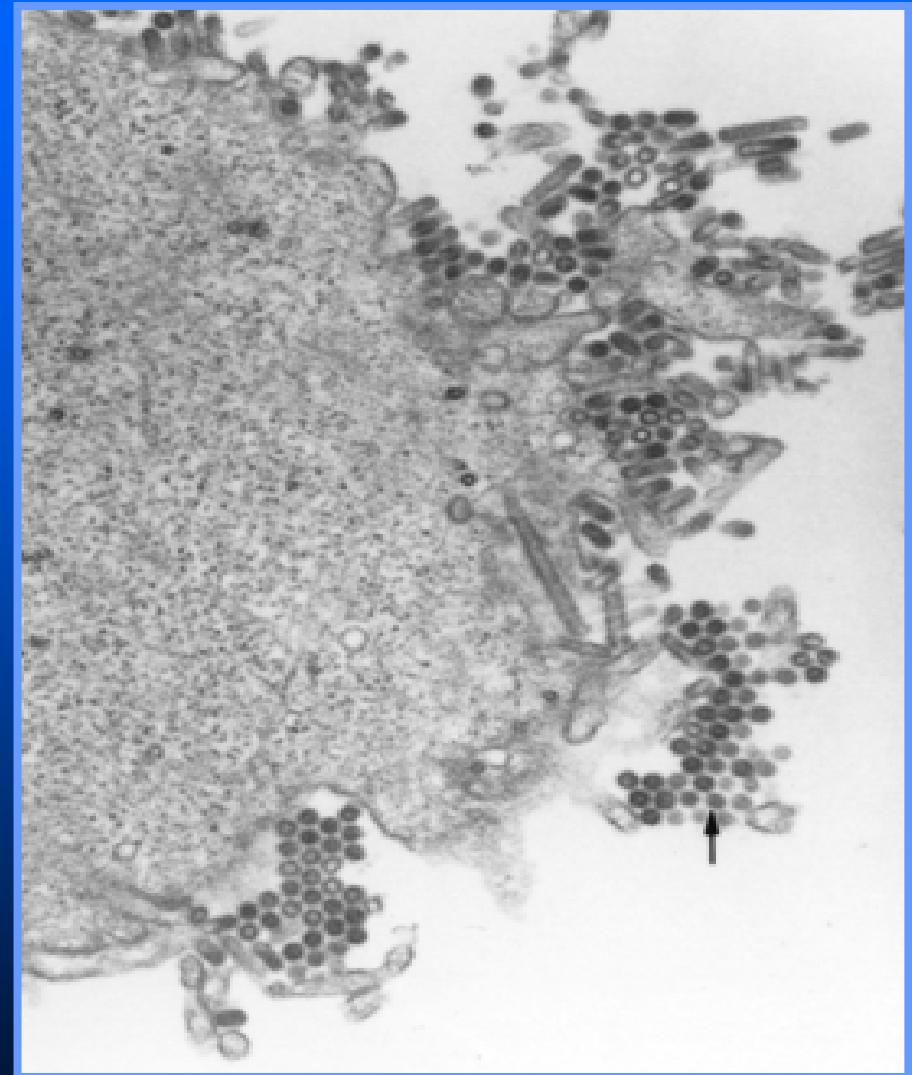
INFLUENZAVIRUS REPLICATIE



NEURAMINIDASE INHIBITIE



MDCK cells + influenza A



MDCK cells + influenza A
+ ZANAMIVIR