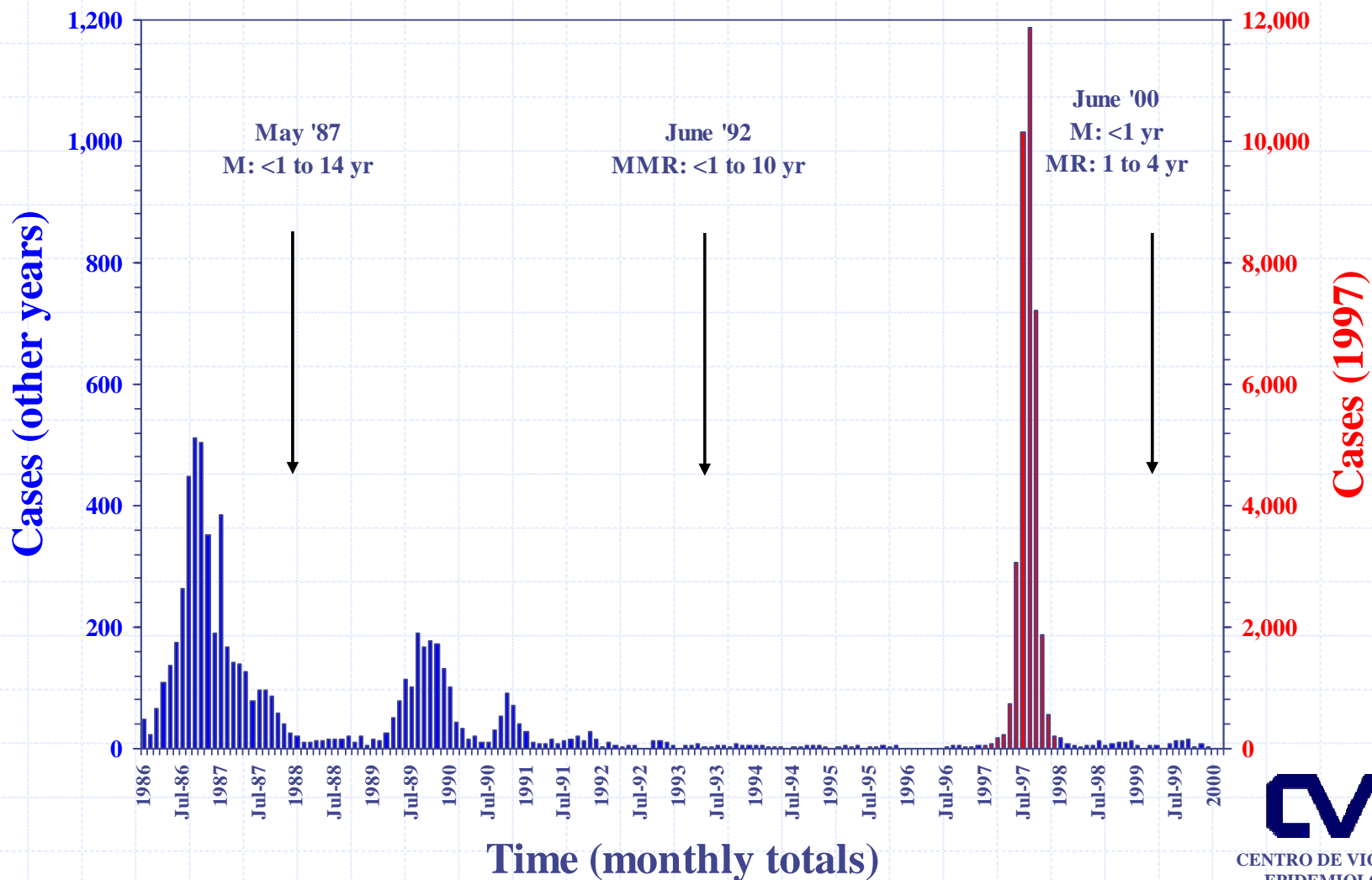


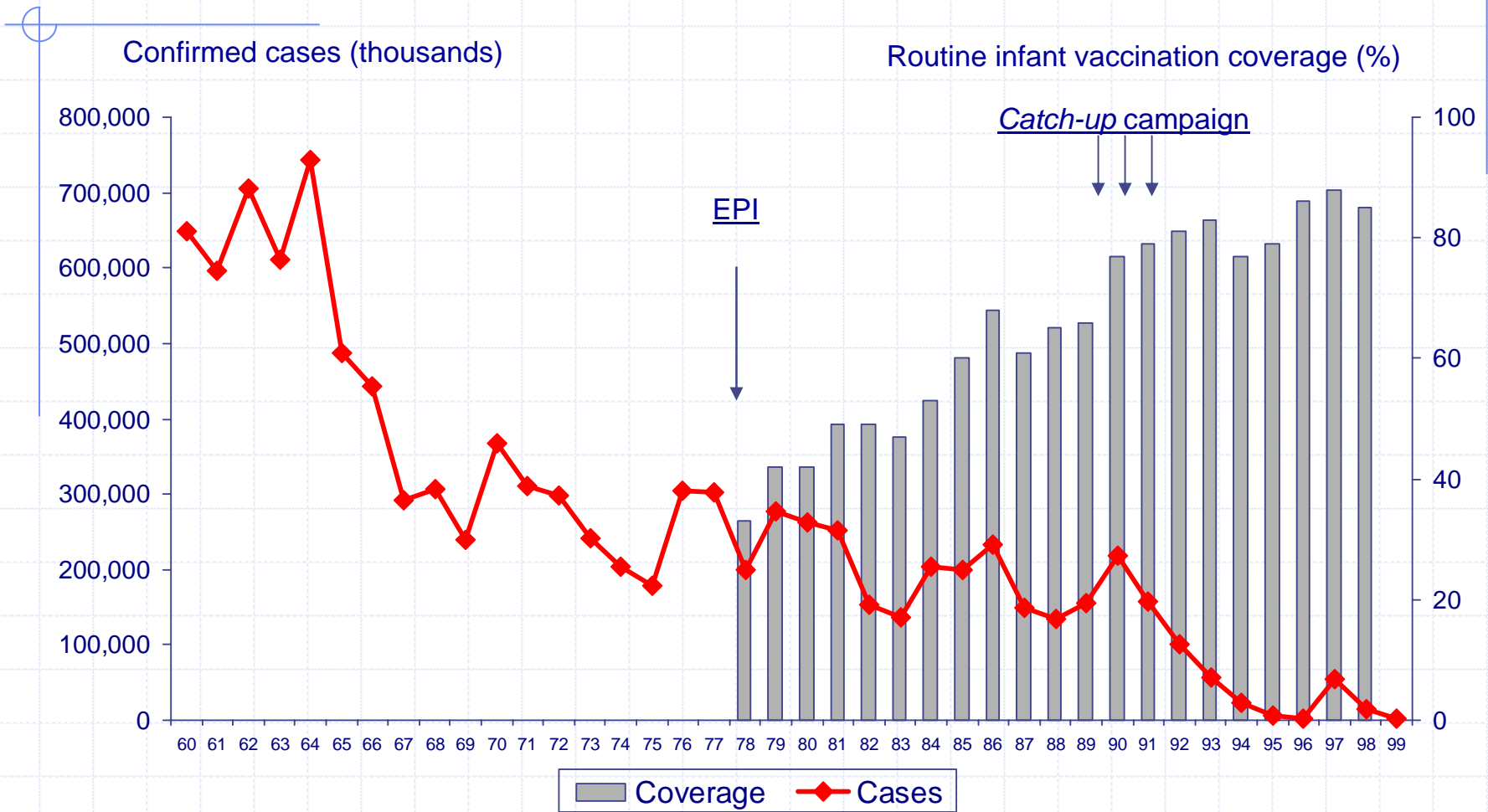
1997 Measles Outbreak in São Paulo, Brazil: Evaluation of Vaccination Strategies via Mathematical Modeling

Secretaria de Estado da Saúde de São Paulo;
Faculdade de Medicina da Universidade de São Paulo; Centers for Disease Control and Prevention; Pan American Health Organization

Measles in Metropolitan São Paulo, 1986-'99



Vaccination Coverage and Reported Cases of Measles, Region of the Americas, 1960-1999*#



Source: PAHO/WHO

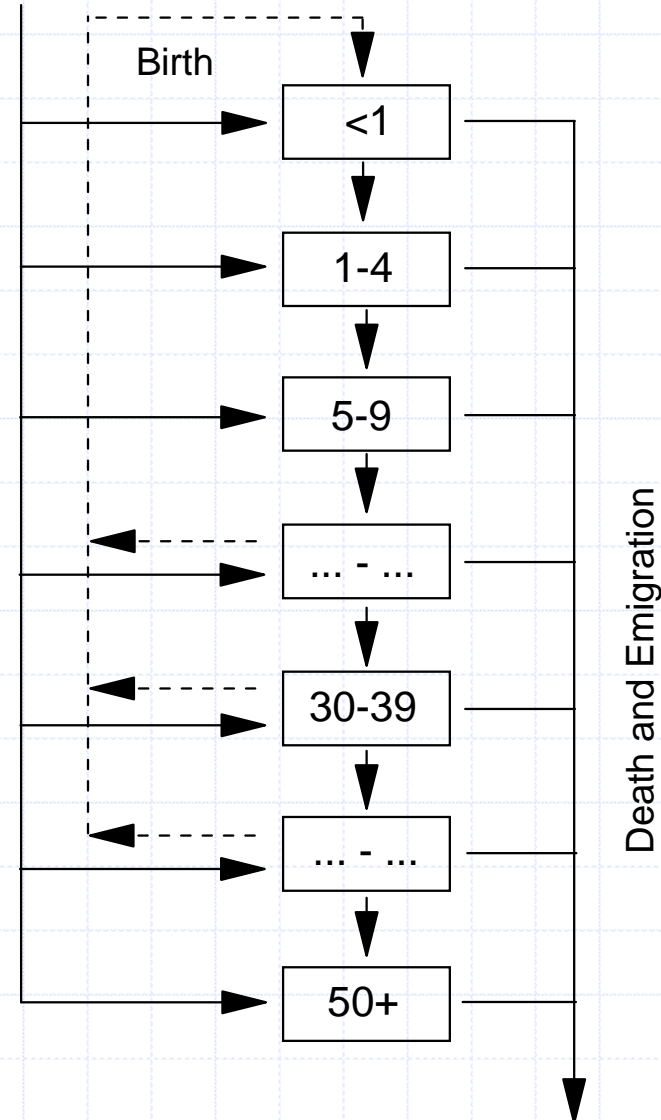
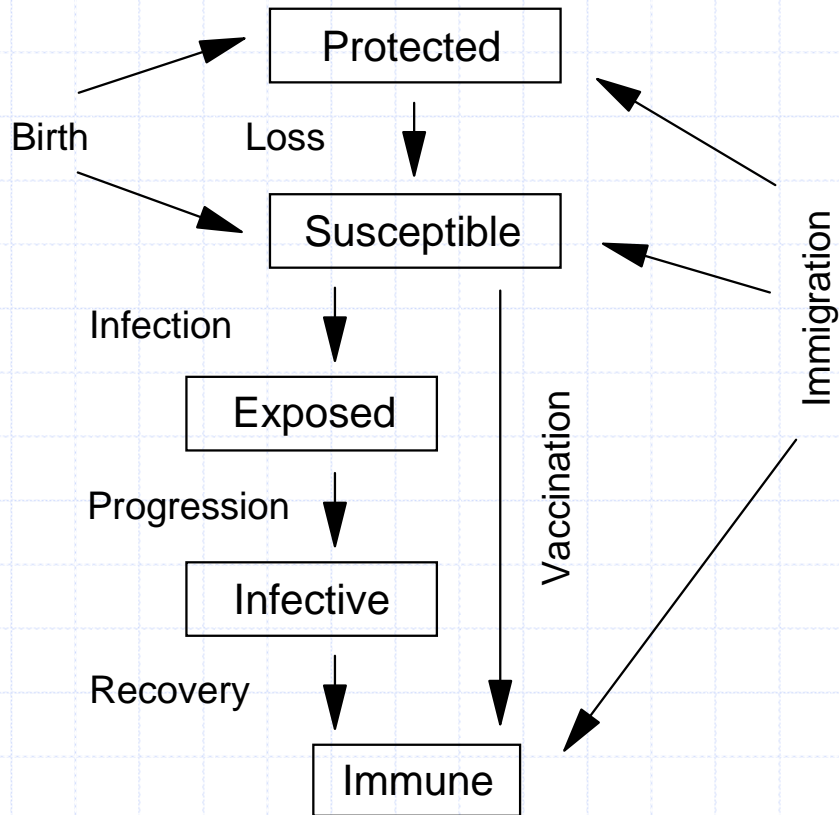
* Data as of 16 October 1999 -2,372 confirmed cases

Coverage data for children <1 year of age
1999 coverage not yet available

Questions

- Would a follow-up campaign during June-August of 1996 have prevented or ameliorated the '97 outbreak?
- What impact did efforts to control this outbreak have?
- Would vaccination of immigrants, particularly from northeastern states, reduce the risk of future urban outbreaks?

Conceptual Model I



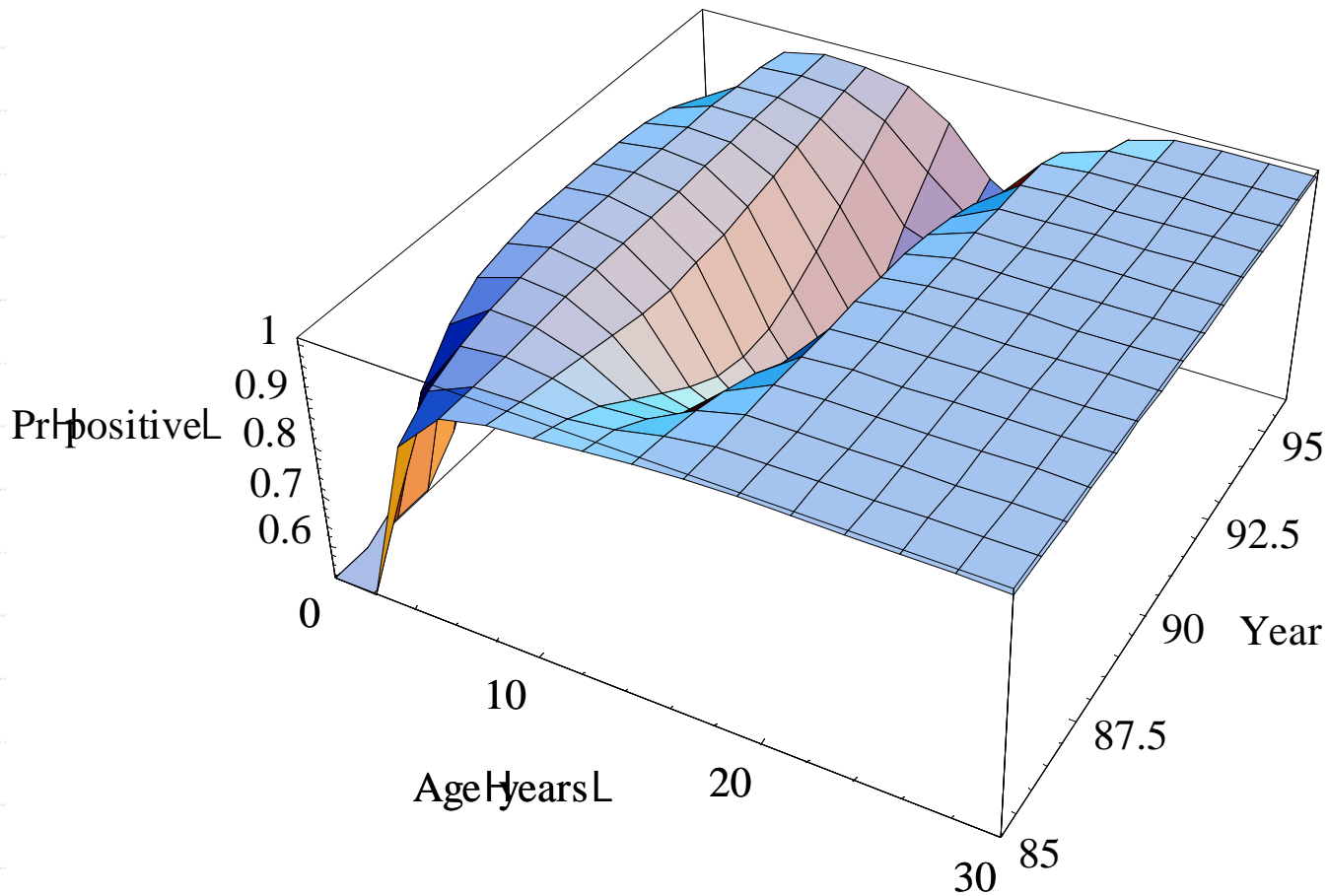
Parameters

- **Demographic:** Vital statistics, including not only resident population and births and deaths, but immigration and emigration rates
- **Programmatic:** Age-specific vaccine efficacy, age distributions of routine doses, timing and target ages of campaigns, coverage
- **Epidemiologic:** Proportions of each resident age class susceptible/immune, characteristics of immigrants, natural history of disease

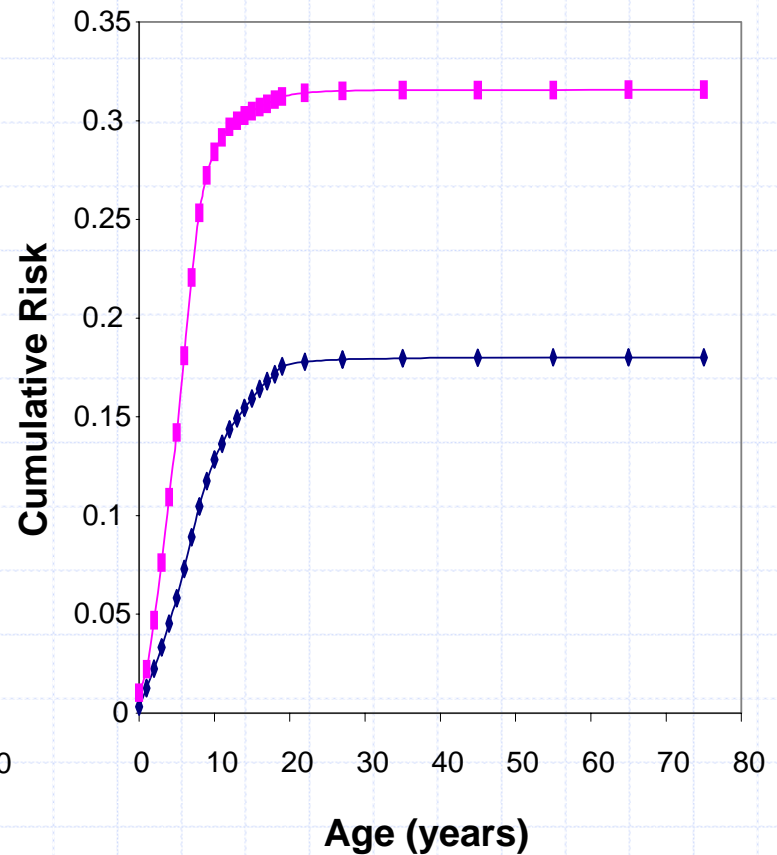
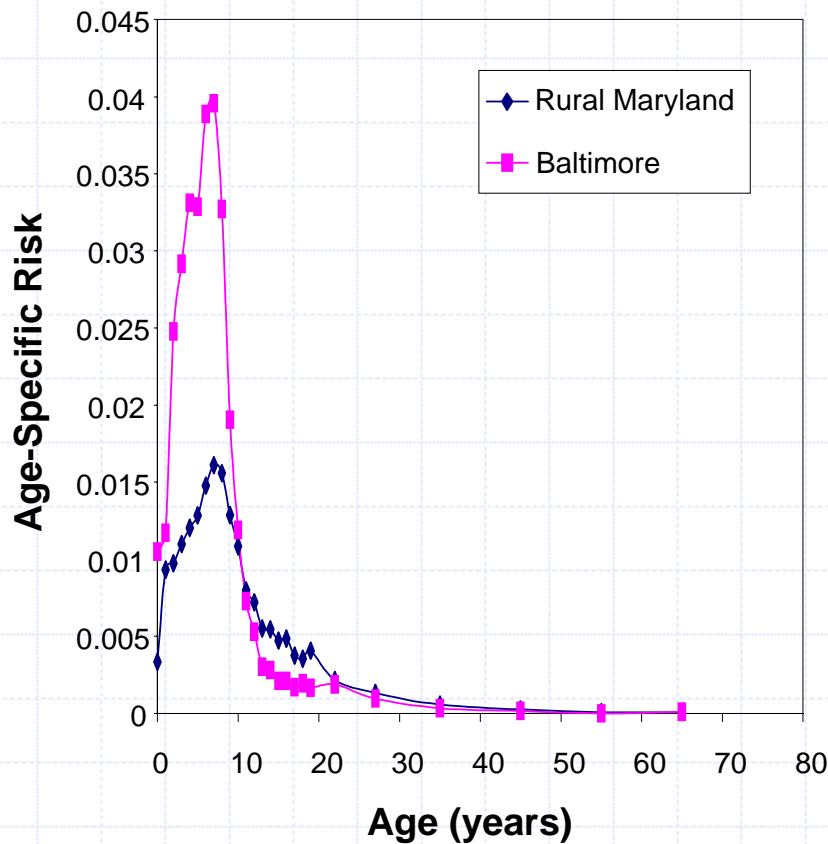
Approach

- Setting-specific information supplemented from elsewhere, informed opinions
- Several parameters adjusted to increase concordance between simulated and reported cases via direct-search algorithms
- Deduced impact of actual and hypothetical interventions, conditional on routine childhood vaccination, experimentally
- Model of loosely-coupled rural, urban and recent immigrant sub-populations planned

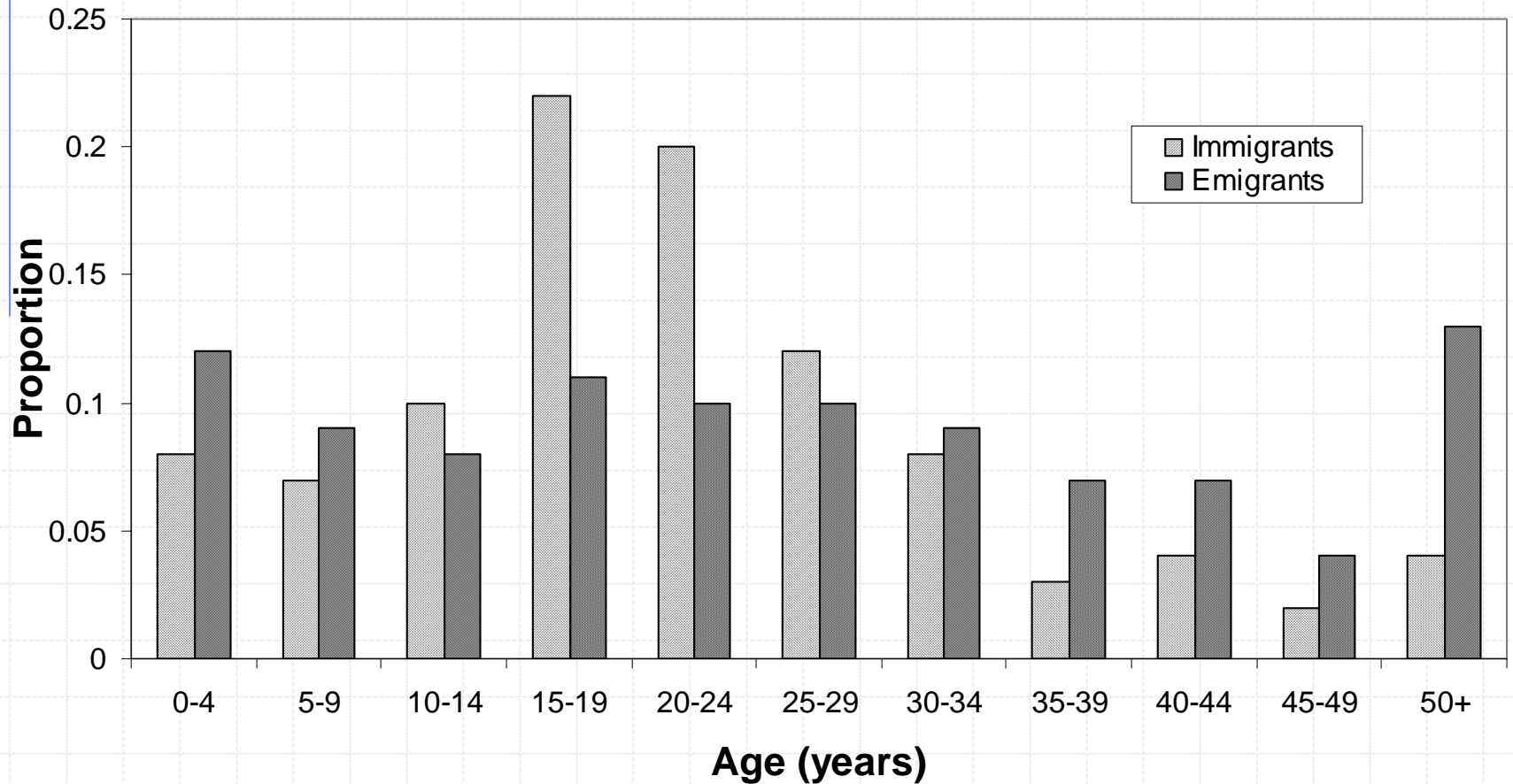
Measles Immunity in São Paulo



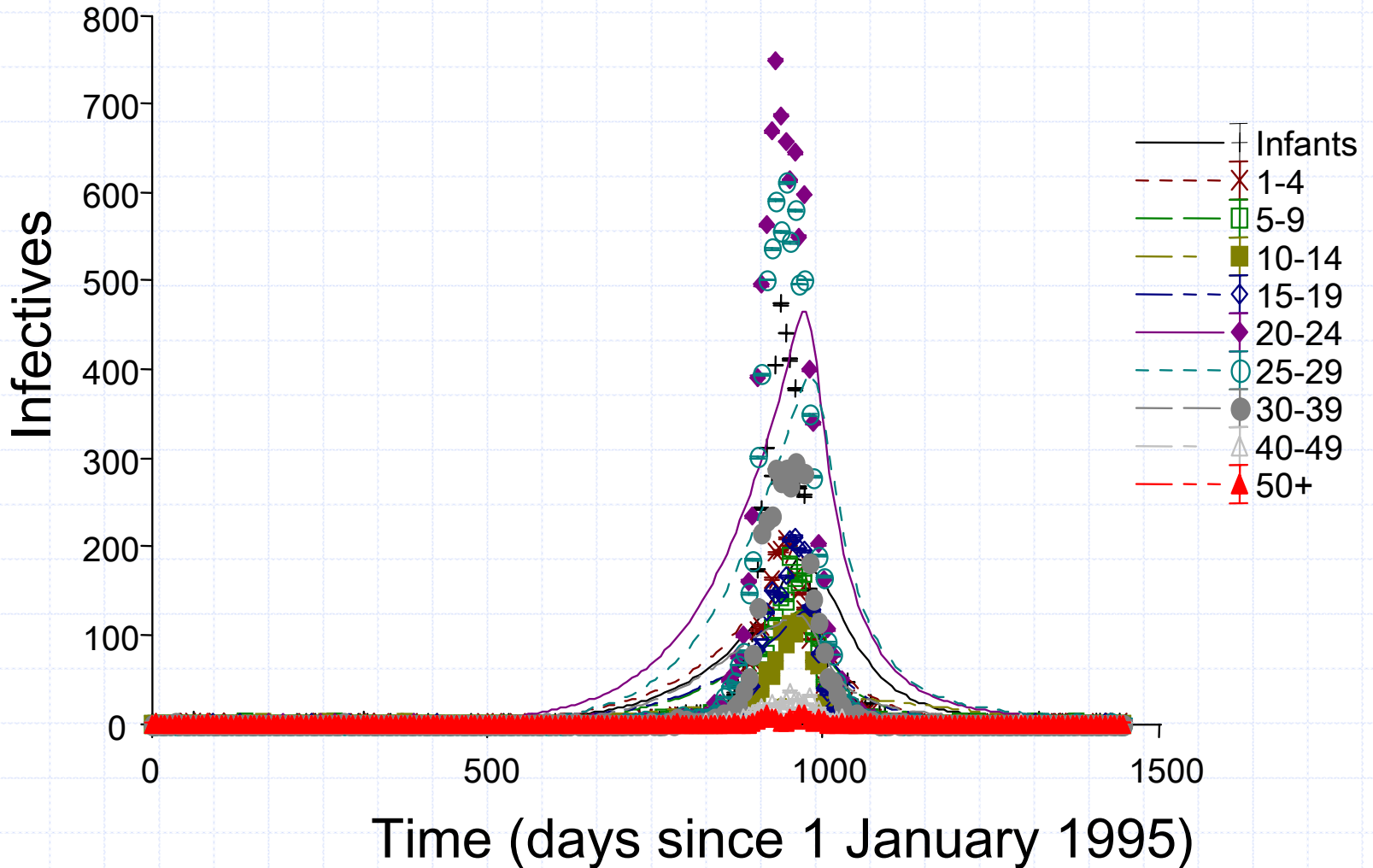
Forces of Infection from Measles Histories in Maryland, 1908-'17



Migration among Residents of São Paulo for <4 Years, 1994



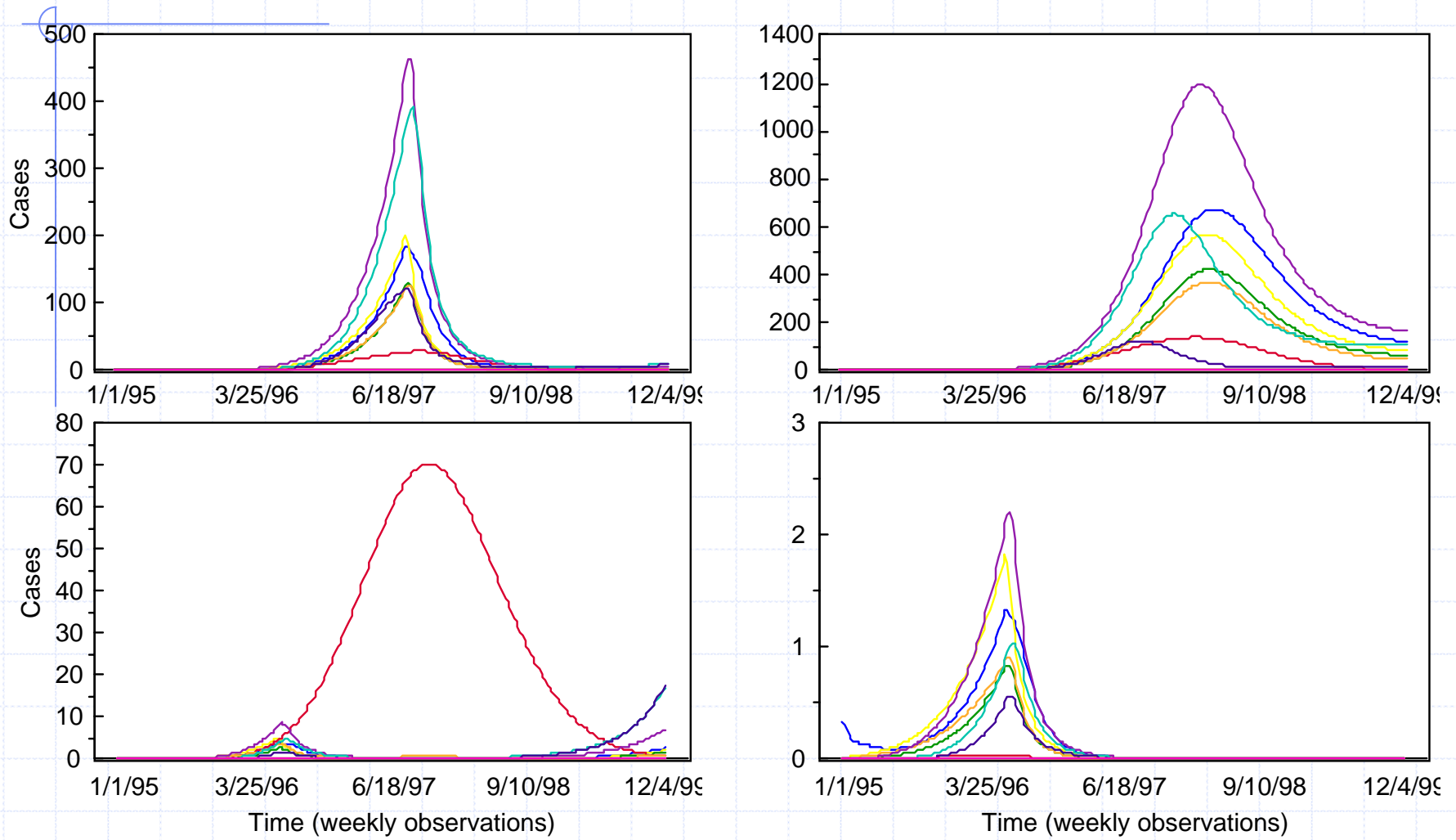
São Paulo, Brazil



Three Experiments: Compare Successive Pairs of Four Scenarios

- With routine two-dose childhood program and outbreak-control efforts
- Without outbreak-control efforts
- With a follow-up campaign during the week of 15 June 1996
- With supplementary adult vaccination having the age distribution of immigrants

Simulated Actual and Three Hypothetical Scenarios



Qualitative Results

- Control efforts directed solely at children were surprisingly effective, given that most cases were over 20 years of age
- Follow-up campaign would have been even more effective, but missed susceptible adolescents, permitting future outbreaks
- Vaccination of young adults, possibly via workplaces of unskilled recent immigrants, would prevent outbreaks

Limitations and Refinements

- More reliable current and historical coverage estimates
- Representative historical or immediately *pre-outbreak* serological surveys
- Historical and projected age-specific rates of immigration and emigration
- Serosurveys where immigrants originate to determine age-specific immunity

Implications of Conclusions

- Some susceptible young adults were long-time urban residents, neither vaccinated as children nor infected subsequently
- But most recently immigrated, by virtue of disparate economic opportunities, from areas whose population immunity is less
- Insofar as neither phenomenon is unique to São Paulo, policymakers must consider heterogeneity within and among populations

Postscript

- Adolescent/young adult immigrants not targeted as recommended, but ...
- A mass campaign was conducted June 2000 to avert predicted measles outbreak
- An outbreak of rubella with similar age distribution occurred soon afterwards in São Paulo, ...
- With, consequently, many cases of congenital rubella syndrome[†], ...
- That, in turn, were followed by a rubella catch-up campaign among women 20-29 years of age

[†]Lanziere et al. 2003. *Pediatr Infect Dis J* 22:323-29;
2004. *Pediatr Infect Dis J* 23:1116-22