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Mortality and morbidity estimates for influenza infection in Asia-Pacific countries

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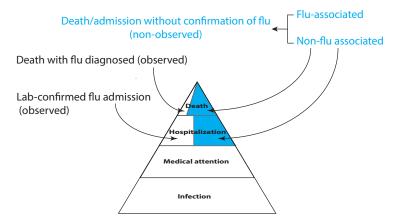
11 June 2015



Introduction

- Influenza viruses cause infections and diseases in all age groups during epidemics.
- Severe disease particularly in the extremes of age.
- Incidence of influenza virus infection varies as high as 20-30% in children and 10-20% in adults in epidemics. Most infections associated with mild disease, some fraction of infections are asymptomatic.

Background



Two general approaches to assess burden at the top of the pyramid in specific populations: (1) laboratory confirmation of influenza in severe cases from a population; (2) infer burden from statistical analysis of administrative data.

Background

- Brooks et al. 2010 PIDJ
- Studied a defined population of children <5y in the catchment area of the study site, for 3.75 years
- Found an overall rate of 511 pneumonia episodes/1000 child-years
- Sampled 12,000 children with respiratory and febrile illness syndromes, estimating that the incidence of influenza-specific pneumonia was 28.6 episodes per 1,000 child-years



Table: Average annual hospitalization rates (per 10,000 population per year) associated with influenza A and B by age in Hong Kong, 2004-2013.

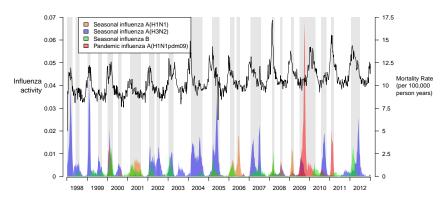
| | Average hospitalization rate (per 10,000 population per year) | | | | | | | |
|--------|---|--------------|-------|--------------|-------|---------------|------|--------------|
| Age | sH1N1 | (95% CI) | sH3N2 | (95% CI) | pH1N1 | (95% CI) | В | (95% CI) |
| 0-6m | 16.9 | (4.3, 40.8) | 31.7 | (15.8, 56.7) | 63.6 | (29.1, 120.8) | 5.8 | (0.7, 20.8) |
| 7-12m | 19.9 | (6.5, 46.5) | 37.4 | (19.9, 64.0) | 56.5 | (24.4, 111.4) | 11.5 | (3.1, 29.5) |
| 2-4y | 21.0 | (10.1, 38.6) | 35.2 | (22.3, 52.8) | 61.5 | (35.2, 99.9) | 13.8 | (6.3, 26.2) |
| 5-9y | 19.5 | (13.1, 27.8) | 36.0 | (28.3, 45.1) | 33.5 | (22.1, 48.7) | 25.9 | (19.5, 33.8) |
| 10-14y | 7.4 | (4.7, 11.0) | 8.1 | (5.6, 11.3) | 19.0 | (12.3, 28.1) | 13.9 | (10.6, 18.0) |
| 15-17y | 1.2 | (0.4, 2.9) | 1.8 | (0.8, 3.3) | 7.0 | (3.6, 12.2) | 3.3 | (1.9, 5.3) |

Based on sampling once per week from two hospitals with a defined catchment population.

Influenza A: Chiu et al 2014 PLoS One; Influenza B: unpublished data



Influenza activity and excess mortality in HK



Estimation of excess influenza-associated mortality using regression models of the form: average weekly death rate = beta * weekly influenza activity $+ \dots$

Annual excess influenza-associated mortality rates in HK

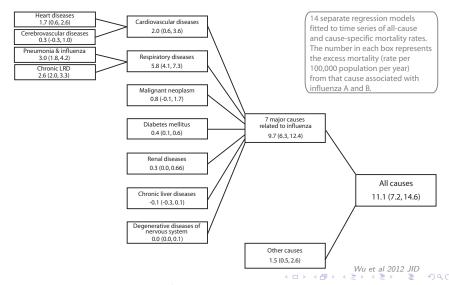
Table: Average type/subtype-specific annual excess all-cause mortality rates by age in Hong Kong, 1998-2009.

| | Average excess mortality rate (per 100,000 population per year) | | | | | | | | | |
|--------|---|--------------|-------|--------------|-------|---------------|------|-------------|------|---------------|
| Age | sH1N1 | (95% CI) | sH3N2 | (95% CI) | pH1N1 | (95% CI) | В | (95% CI) | All | (95% CI) |
| 0-4y | 0.9 | (-0.3, 2.4) | 0.2 | (-1.7, 2.2) | -3.0 | (-7.6, 2.5) | -1.5 | (-4.0, 0.8) | -0.7 | (-3.7, 2.4) |
| 5-14y | 0.1 | (-0.2, 0.5) | -0.1 | (-0.5, 0.4) | -0.7 | (-1.9, 0.7) | -0.3 | (-0.8, 0.3) | -0.2 | (-1.0, 0.5) |
| 15-44y | 0.8 | (0.1, 1.3) | 0.6 | (0.0, 1.4) | -0.8 | (-2.6, 1.0) | 0.0 | (-0.8, 0.9) | 1.3 | (0.3, 2.4) |
| 45-64y | 0.2 | (-1.5, 1.8) | 1.9 | (-0.1, 3.9) | -0.8 | (-6.4, 5.3) | 1.3 | (-1.3, 3.6) | 3.3 | (0.1, 6.1) |
| ≥65y | 8.4 | (-6.0, 21.7) | 58.8 | (40.5, 76.0) | 23.4 | (-23.7, 75.7) | 20.3 | (0.1, 41.5) | 89.7 | (61.8, 113.7) |
| All | 1.6 | (-0.3, 3.3) | 6.9 | (4.3, 9.4) | 2.2 | (-4.8, 8.8) | 2.5 | (-0.5, 5.3) | 11.1 | (7.2, 14.6) |

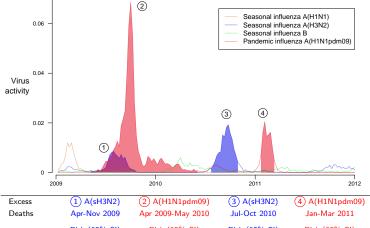
Wu et al 2012 JID



Excess mortality by cause



Second wave of pandemic influenza in Hong Kong



| Excess | (1) A(sH3N2) | (2) A(H1N1pdm09) | (3) A(sH3N2) | (4) A(H1N1pdm09) |
|-------------|----------------|-------------------|----------------|------------------|
| Deaths | Apr-Nov 2009 | Apr 2009-May 2010 | Jul-Oct 2010 | Jan-Mar 2011 |
| | Risk (95% CI) | Risk (95% CI) | Risk (95% CI) | Risk (95% CI) |
| All-cause | 3.5 (0.3, 6.8) | 2.1 (-8.1, 11.8) | 6.2 (3.3, 9.0) | 4.7 (-0.7, 10.5) |
| Respiratory | 1.7 (0.2, 3.0) | 1.2 (-3.2, 5.5) | 2.9 (1.5, 4.0) | 3.9 (1.7, 6.4) |



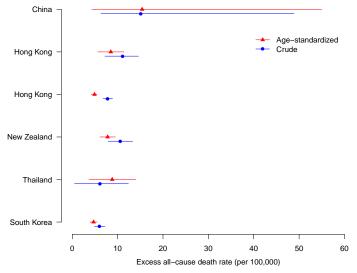
Influenza-associated mortality, selected locations

Mortality 000000

Table: Average annual excess all-cause mortality rates in all ages in selected Asia-Pacific locations.

| Location | Period | Rate | (95% CI) | Reference |
|-------------|-----------|------|-------------|----------------------------|
| China | 2004-2009 | 15.1 | (6.3, 48.9) | Yu 2013 IORV |
| Hong Kong | 1998-2009 | 11.1 | (7.2, 14.6) | Wu 2012 JID |
| Hong Kong | 1998-2009 | 7.8 | (6.8, 8.9) | Yang 2012 Epidemiol Infect |
| New Zealand | 1990-2008 | 10.6 | (7.9, 13.3) | Kessaram 2014 IORV |
| South Korea | 2003-2013 | 6.0 | (4.9, 7.2) | Cowling unpublished data |
| Thailand | 2005-2009 | 6.1 | (0.5, 12.4) | Cooper 2015 AJE |

Accounting for different demographics in different locations



Burden of medically-attended ILI, Mongolia

Table 3. Number of ILI cases and incidence per 1000 population in Selenghe during the two influenza epidemic periods

| Age group | Week 6, 20 | 9 influenza period 109–week 10, 2009 5 weeks) | 2009–2010 influenza period Week 43, 2009–week 6, 2010 (17 weeks) | | |
|-------------|------------------------|---|--|-------------------------------|--|
| | Number of ILI cases | Incidence per 1000 population | Number of ILI cases | Incidence per 1000 population | |
| 0-11 months | 13 | 3.7 | 78 | 21.9 | |
| 1-4 years | 39 | 2.8 | 157 | 11.3 | |
| 5-9 years | 26 | 1.4 | 108 | 5.6 | |
| 10-14 years | 20 | 1.0 | 82 | 4.2 | |
| 15-24 years | 14 | 0.3 | 112 | 2.4 | |
| 25-44 years | 9 | 0.1 | 92 | 1.3 | |
| 45-64 years | 4 | 0.1 | 22 | 0.6 | |
| ≥ 65 years | 0 | 0.0 | 6 | 0.7 | |
| Total | 125 | 0.6 | 657 | 3.1 | |

Nukiwa et al 2011 WPSAR



Why should influenza be a public health priority?

- The burden of influenza is substantial in Asia-Pacific countries, pneumonia is a leading cause of hospitalization and death in the extremes of age and influenza is an important viral cause of pneumonia which is preventable by vaccination.
- Limitations most data on disease burden come from a small number of locations, more studies would be valuable. not only on the health impact but also the broader socio-economic impact of influenza epidemics.
- Not included in this talk burden and impact of avian influenza



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