Reduction in HPV-Associated High Grade Cervical Lesion Incidence in Connecticut, 2008-14: Evidence for Herd Immunity

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Background

- Annually responsible for ~12,000 cases/yr of cervical cancer and >200,000 cases of high grade cervical cancer precursor lesions (CIN2+)
- CIN2+ (and cervical cancer) became vaccine preventable in 2006
 - HPV vaccine recommended for girls 11-12 yrs in 3-dose series
 - Catch-up for women up to 26 yrs
 - >95% effective in preventing CIN2+ (cervical interepithelial neoplasia grades 2 and higher, adenocarcinoma in situ)

Surveillance - HPV IMPACT

- In 2008, CDC STD staff and five EIP sites (CA, CT, OR, TN, NY) established active, population-based surveillance for CIN2+
 - Pathology-laboratories
 - In CT, CIN 2+ made reportable statewide

Study Objectives

- Determine trends in CIN 2+ incidence in Connecticut, 2008 through 2014
- Compare observed to expected changes in incidence in 21-24 year olds

Methods 1

- Use deduplicated cases assigned to the year in which their initial diagnosis occurred
- Determine age & age-group specific incidence rates using annual population estimates (US Census)
- Compared observed to maximum expected decrease through 2014 in 21-24 year olds
 - 21 is youngest age for routine PAP screening
 - 21-24 year olds in 2014 were 13-16 years old in 2006, group most likely to be HPV-infection naïve when vaccinated.

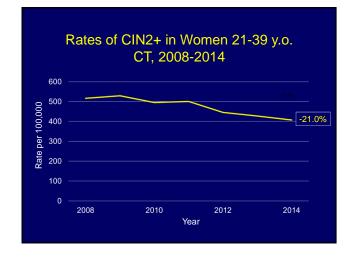
Maximum Expected Decrease 21-24 yo, CT

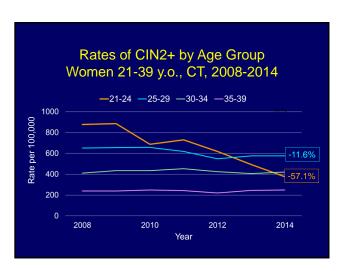
[max. % of CIN2+ that are vaccine-preventable] x [est. max. percentage vaccinated in age group]

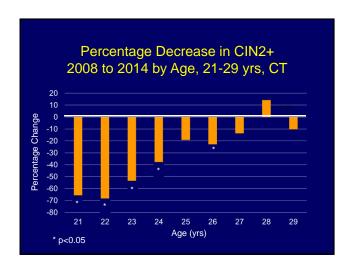
- % CIN2+ vaccine-preventable = [100% types 16/18 + 21% other HPV types] = $54.3\%^* + 9.6\%^{**} = 63.9\%$
- % vaccinated = 66%***

Max expected decrease = 42.2%

- Hariri. Cancer Epidemiol Biomarkers Prev 2015;24:393-399
 Wheeler. Lancet Oncol 2012;13:100-10
 NIS Teen 2008-2011.







Observed vs. Maximum Expected Decrease in CIN2+ in 21-24 year olds [max. % cases potentially preventable: 63.9%] x

[max. % cases potentially preventable: 63.9%] x [max % vaccinated: 66%] = 42.2% (314 cases*)

Actual decrease = 57.1% (425 cases*), p=0.00004

Relative actual to expected decrease: 1.36 [95% CI 1.17-1.57]

* Using 2008 denominator and baseline 744 cases

Hypotheses

- Cross-protection for non-vaccine strains
- Under-reporting in 2014 relative to 2008
- Reduced rates of detection of CIN2+ due to changes in PAP screening recommendations in 2012
- Herd immunity

Conclusions

- There has been a progressive decrease in CIN2+ diagnoses, mostly since 2011.
- Vaccination rates may be high enough in those who were 13-16 years (now 21-24) when vaccine licensed to reduce the probability of HPV exposure in the unvaccinated, leading to herd immunity.

Acknowledgements

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Limitations

- Don't know what HPV types responsible for the changes observed – changes in behavior could account for some of the decrease.
- Percentage of CIN2+ prevented in 21-22 yo already at estimated maximum – suggests some parameters could be off.
- Estimate of % of CIN+ that are vaccine types were from EIP data collected 2008-2012 – possible underestimate of % CIN2+ preventable.

