



Published in final edited form as:

Vaccine. 2011 March 3; 29(11): 2149–2158. doi:10.1016/j.vaccine.2010.12.078.

From the Patient Perspective: the Economic Value of Seasonal and H1N1 Influenza Vaccination

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Abstract

Although studies have suggested that a patient's perceived cost-benefit of a medical intervention could affect his or her utilization of the intervention, the economic value of influenza vaccine from the patient's perspective remains unclear. Therefore, we developed a stochastic decision analytic computer model representing an adult's decision of whether to get vaccinated. Different scenarios explored the impact of the patient being insured versus uninsured, influenza attack rate, vaccine administration costs and vaccination time costs. Results indicated that cost of avoiding influenza was fairly low, with one driver being required vaccination time. To encourage vaccination, decision makers may want to focus on ways to reduce this time, such as vaccinating at work, churches, or other normally frequented locations.

Keywords

Influenza Vaccine; H1N1 Patients; Economics

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1. INTRODUCTION

Vaccination coverage for influenza among adults has remained well below the desired rates set forth in *Healthy People 2010*.^[1–2] Approximately 32% of the general public and less than 70% of high risk groups identified by the Advisory Committee on Immunization Practices (ACIP) reported being vaccinated against influenza during the 2008–2009 flu season.^[1–2] The monovalent 2009 H1N1 influenza vaccine experienced particularly poor uptake, as approximately 80% of the U.S. population remained unvaccinated.^[3] Misinformation and lack of knowledge regarding vaccine safety, effectiveness and importance, as well as challenges of cost, location and time have been frequently identified as barriers to achieving higher coverage.^[4–6]

While studies have quantified the economic value of influenza vaccination from the societal and third party payor perspectives in a variety of patient populations, few have addressed the economic value of vaccination from the perspective of individual patients.^[7–10] Although it is unlikely that patients base their decisions about vaccination purely on economic considerations, better understanding a vaccine's economic value to patients can assist various types of decision-making. Studies have demonstrated that a patient's perceived cost-benefit of a medical intervention can affect his or her utilization of the intervention, i.e., medical intervention utilization is fairly price elastic.^[11–12] For example, pharmacy copayment increases have been paralleled by reductions in patient use of psychiatric and cardiovascular medications.^[13–14] Conversely, decreases in required copayments have been followed by increased utilization of antibiotics.^[15–19] A recent survey suggested that higher vaccine cost may dissuade adults from getting vaccinated.^[12] As a result, the Task Force on Community Preventive Services recommends reducing client out-of-pocket costs as a means of enhancing access to vaccination services and increasing vaccination coverage.^[20]

Therefore, better understanding the costs of vaccination for a patient could be useful for various decision makers associated with vaccination efforts. Health care workers and public health officials may use the information to convey the advantages and disadvantages of vaccination to patients. It could also serve as the basis for vaccine pricing, third party coverage, incentive, and various cost-sharing decisions. For instance, identifying disparities in the economic value for under- or uninsured patients may motivate programs to alleviate their financial burden in getting vaccinated. Value-based insurance design, the idea of promoting the use of a service (such as preventive influenza vaccination)—through lower or no cost immunization—when the clinical benefits exceed its cost, is an approach that may help improve vaccine coverage. Ascertaining the drivers of patient cost can help to identify operations structures and policies amenable to intervention.

To delineate the economic value of influenza vaccination from the adult (between 18 and 49 years old) patient perspective, we developed a computer simulation model representing the decision to get vaccinated. Different scenarios explored the impact of the patient being insured versus uninsured, influenza attack rate, and vaccine administration costs and wait times.

2. METHODS

2.1 Model Structure

Using TreeAge Pro 2009 (TreeAge Software, Williamstown, Massachusetts), we developed a decision analytic computer simulation model representing an individual's decision of whether to get vaccinated against influenza. The model focused on standard-risk adults (i.e., not high-risk by ACIP definitions), who comprise the majority of individuals in the 18 to 49

year old population and are eligible for both the TIV and LAIV vaccines. Outcomes data for the models came from an extensive literature review, using the following search terms: [seasonal AND influenza], [2009 H1N1], [influenza AND vaccination], [influenza AND vaccination AND cost], [influenza AND outcomes], and [influenza AND vaccination AND side effects]. The authors judged each article critically on study design and relevance to our target population. Separate scenarios assumed that the patient had commercial insurance or was uninsured. Additional scenarios examined the use of tetravalent inactivated vaccine (TIV) versus live attenuated influenza vaccine (LAIV). Figures 1 and 2 depict the general structure and influenza outcomes subtree of the model, respectively, which assumed a patient perspective.

Patients who received the vaccine then had probabilities of experiencing major or minor side effects or both. Minor side effects included local inflammation or minor systemic flu-like symptoms, requiring self-treatment with ibuprofen. Guillian-Barré Syndrome (GBS), an autoimmune disorder of the peripheral nervous system, was the only major side effect considered; all cost and probability data relating to major side effects from vaccination in the model were therefore GBS- specific. The risk of developing symptomatic influenza depended on the clinical attack rate and whether the patient received a vaccine. Vaccination attenuated the risk of influenza by an amount determined by vaccine efficacy. A patient who developed symptomatic influenza then proceeded through the influenza outcomes sub-tree (Figure 2). A symptomatic patient had a probability of visiting a medical clinic, being hospitalized, requiring admission to the intensive care unit (ICU), and not surviving. Symptomatic patients that did not interface with the health care system relied on over-the-counter medications. Of the patients who visited a clinic, half (50%) received antiviral medication treatment.

Patient costs originated from three sources:

- **Productivity/Wage Loss:** Getting vaccinated, contracting symptomatic influenza, and hospitalization each incurred unique durations of time off from work. Productivity losses were equivalent to the wages that the patient could have earned during the required time off from work.
- **Medication/Treatment Costs:** Vaccines cost the patient money. Symptomatic patients purchased over-the-counter medications. Half of all patients who visited the clinic paid for prescription antivirals. Prescription medication and vaccine costs depended on the patient's insurance status
- **Health Care Costs:** Patient payments for clinic visits, hospitalization, and ICU stays depended on the patient's insurance status.

For each simulation run, the equation below computed the cost per influenza case averted by the vaccine:

$$=(\text{Cost}_{\text{Vaccination}} - \text{Cost}_{\text{No Vaccination}}) \div (\text{Number of Influenza Cases}_{\text{without Vaccination}} - \text{Number of Influenza Cases}_{\text{with Vaccination}}) \quad (1)$$

The following equation calculated the cost per influenza death averted by the vaccine:

$$=(\text{Cost}_{\text{Vaccination}} - \text{Cost}_{\text{No Vaccination}}) \div (\text{Number of Deaths}_{\text{without Vaccination}} - \text{Number of Deaths}_{\text{with Vaccination}}) \quad (2)$$

2.2 Data Inputs

Table 1 organizes the data inputs for our model into costs, probabilities, and relevant time durations and lists their ranges and distributions. Cost and efficacy parameters were specific

to the type of vaccine. Table 1 also lists the sources of each parameter input. The amounts paid by each individual depended on his or her insurance type, where uninsured patients had to cover medical costs (not charges).[21] Hospitalization costs came from the healthcare utilization project's (HCUP) national inpatient survey (state inpatient data from Arizona, Florida, Massachusetts, Maryland, New Jersey, New York, Texas and Washington), using ICD-9 codes 480 and 357 for hospitalization and ICU admittance from influenza and major side effects such as Guillain-Barré respectively. Clinic costs originated from the Centers for Medicare and Medicaid Services (CMS). Medication costs were derived from their average wholesale prices, as drawn from the PDR Red Book.[22] The 2008 national median wage, retrieved from the Bureau of Labor and Statistics, was used to create a distribution of daily wages.[23]

By contrast, the model made the following assumptions about the commercially insured based on benefit design information obtained from the Henry J. Kaiser Family Foundation, Health Research & Educational Trust Survey of Employer-Sponsored Health Benefits, 2009:

- 36.3% had no annual deductible [24]
- The distribution of plan types was as follows: 20% health maintenance organization (HMO), 60% preferred provider organization (PPO), 10% point of service (POS), and 10% high deductible health plan (HDHP/SO) [24]
- 18% of those with no deductible had coinsurance [24]
- Those who paid a deductible (63.7%) paid \$809.90. [24]

Based on a report using data collected through the Medical Expenditure Panel Surveys, uninsured individuals paid approximately one third of their total medical costs. [25] All costs were expressed in 2009 U.S. dollars with a 3% discount rate converting costs from other years.

2.3 Sensitivity Analyses

Probabilistic (first and second-order) sensitivity analyses with microsimulation simultaneously varied the values of each variable across the ranges listed in Table 1, i.e., each parameter value from distributions listed in Table 1 and then individuals proceeded down each branch of the tree based on stochastic dice rolls against the probability values drawn from the distributions. Specific analyses explored the effects of ranging vaccination cost, which included both the cost of vaccine and other ancillary costs (i.e., travel and/or childcare expenses). We utilized a baseline cost of \$15 (TIV) and \$25 (LAIV) for scenarios where vaccination costs were equivalent to only that the vaccine itself as well as \$0, \$50, and \$150 for both TIV and LAIV. The \$0 vaccination cost accounts for scenarios where vaccine may be administered during a routine visit and does not incur an additional cost; higher vaccine costs (\$50 and \$150) allow for the inclusion of potential transportation costs needed to get to a health care facility. We additionally varied the hours missed from work to get vaccinated (1 hour to 4 hours), the probability of major vaccine side effects (i.e., Guillain-Barré), the clinical attack rate of influenza, and the probability of influenza resulting in a hospital and ICU admission.

Baseline attack rates for seasonal influenza drew from a beta distribution (estimated from a mean of 6.6% and standard deviation of 1.7%). One-way sensitivity analyses also evaluated ranging this attack rate from flat values of 2% up to 15%.[26–27] As limited U.S based data exists regarding the incidence of 2009 H1N1 during the 2009–2010 influenza season, the H1N1 baseline attack rate for the US was estimated using a seroprevalence reports conducted in England during this time period; additional simulations utilized an attack rate of 6.5%, which is comparable to incidence given in a brief U.S. based report.[28–29] While

baseline influenza risks were drawn from studies reporting data from a limited number of influenza seasons, these distributions of influenza risk are likely the most probable. Varying the attack rate higher and lower than these expected distributions was not meant to represent a likely risk of flu, but was instead intended to quantitatively illustrate the effect of an increased or decreased risk of influenza on the cost of avoiding a case of influenza by vaccination. For seasonal flu, the probability of ICU admittance ranged from 5% to 20%; the probability of ICU care being received as a result of H1N1 ranged from 15% to 25%. Two case fatality rates for H1N1 were evaluated; the first distribution (triangular, likeliest value: 0.16% range: 0.066–0.333%) was based on the symptomatic influenza-specific case fatality, while the second was a case fatality rate (triangular, likeliest value: 0.01% range: 0.007–0.016%) calculated out of the overall self-reported symptomatic influenza-like illness reports.[30] Baseline runs used 0.002% and 0.0003% for the probability of major side effects from LAIV and TIV vaccines respectively.[31–32] Additional analyses examined the effects of doubling the probability of ICU admittance and major side effects.

3. RESULTS

3.1 Simulation Parameters

The median age used in the model was 30 years for uninsured patients and 40 years for commercially insured patients. Additional analyses evaluated ages 40 and 49 for uninsured and commercially insured populations, respectively, as 49 is the oldest age at which administering LAIV is still recommended. Median age data was generated with data from the Bureau of Labor and Statistics and a recent paper on the cost and coverage of healthcare among the uninsured.[25,33] Each simulation run sent 1,000 hypothetical individuals, each with a unique set of characteristics (each individual drew from the wage distribution to represent the diversity of jobs, geographic locations, and sociodemographic characteristics that may affect wages; those who developed side effects drew from the cost of side effects distribution and those who became ill drew from the clinic and hospitalization cost distributions to reflect the sociodemographic and clinical diversity that exists among adults), through the model 1,000 times for a total of 1,000,000 patient outcomes for each simulation run. (Note: the parameters drew independently from each respective distribution and were not correlated.)

3.2 Measurements of Stochasticity

Tornado diagrams in Figure 3 demonstrate the impact that each variable included in our sensitivity analysis has on the incremental cost per averted case. As shown in the figure, ranging vaccination cost, the time needed for vaccination, and attack rate had the most impact on the incremental cost for all scenarios, with vaccination cost having the greatest effect; the probability of ICU or mortality from influenza and the probability of major vaccine side effects had little to no impact on the cost averted by preventing an influenza case.

Scatter plots (see Supplementary Figure 1) depict the variability of cost and influenza occurrence within a particular simulation. While vaccinating was associated with higher cost compared to no vaccination, the vaccination group had substantially fewer influenza cases as well as lower variation in influenza occurrence. Relatively little difference in cost and influenza cases was observed between the commercial insurance and uninsured groups for both H1N1 and seasonal influenza. Seasonal influenza scenarios illustrated a lower variation in the occurrence of influenza in the no vaccination group (but not in the vaccination group), which is likely a product of a narrower attack rate baseline distribution (see Table 1).

3.3 Seasonal Influenza Scenario

Table 2 presents the median and 95% confidence interval (both 2.5th and 97.5th percentiles) for the cost per averted influenza case from our seasonal influenza model assuming a 15% baseline probability of ICU (given hospitalization) as well as a 0.0003% and 0.002% probability of major side effects from TIV and LAIV vaccine formulations, respectively. As mentioned above, the value of vaccination was fairly sensitive to vaccination cost, the time required for vaccination, and influenza attack rate. When vaccination took no more than one hour, TIV generated marginal (when compared to no vaccination) costs only slightly higher than that of the vaccine itself (approximately \$27 for seasonal influenza vaccine and \$17 for H1N1 vaccine) shown in Figure 4.

A 4 hour vaccination administration time, which would correspond to a patient traveling to a physician's clinic, was almost three times as expensive as a one-hour administration time, which may correspond to a patient traveling to either a closer location or a location that has less of a wait time (e.g., a dedicated immunization clinic). Although varying vaccination costs over the range explored in the sensitivity analyses had the greatest effect on the economic benefit of vaccination, a slight increase in influenza attack rate results in a greater impact on vaccination benefit. For example, while increasing attack rate from baseline to 15% (approximately 2 times the mean at baseline) resulted in a 3 fold decrease in the cost of averting an influenza case, increasing vaccination cost by the same amount (from \$25 to \$50) resulted in a 2 fold increase in cost per averted case (see Table 2).

TIV consistently provided greater economic value than LAIV, as the added efficacy of LAIV did not overcome the higher cost of the vaccine. The economic value of vaccination was fairly comparable for insured and uninsured patients, mainly because influenza hospitalization was relatively infrequent among adults and insurance status had little impact on the cost of over-the-counter medications and missing work.

3.4 2009 H1N1 Influenza

H1N1 vaccination provided relatively more economic benefit than seasonal influenza vaccination, mainly because H1N1 had slightly higher morbidity and mortality than seasonal influenza among the younger adult population (i.e., the population that comprises much of the commercially insured and uninsured). Table 3 demonstrates the effect of vaccine cost, administration time, and attack rate on the median cost of avoiding an influenza case, where baseline probability of H1N1 influenza was a triangular distribution (min=6.5%, likeliest=11.5%, max=19.4%) ICU from H1N1 (given hospitalization) = 20%, case fatality rate was a triangular distribution (min= 0.007%, likeliest= 0.01%, max= 0.016%), and the probabilities of major side effects from TIV and LAIV vaccine formulations were 0.0003% and 0.002%, respectively. While the economic benefit of vaccination was comparably dependent on both the time off from work needed for vaccination and influenza attack rate, varying vaccine cost over the range evaluated through sensitivity analyses had the greatest effect on the cost averted by vaccination. However, as with seasonal influenza vaccination, a greater impact results with decreased influenza risk than with increased vaccination cost of similar magnitude. No significant difference was present between insured and uninsured individuals. Hospitalization rates for H1N1 influenza were still not high enough to result in differences by insurance status. Additionally, there was little difference between the LAIV and TIV vaccines even if the individual were responsible for higher total vaccination costs (vaccine costs for the 2009 H1N1 vaccination program were fully subsidized by the federal government).

4. DISCUSSION

Compared to the number of published studies from the societal and third party payor perspectives, relatively few economic studies have looked at medical interventions from the patient perspective. The vast majority of health economic studies focus on the societal perspective, summing costs across all key stakeholders, or have evaluated whether a large policy-making entity (e.g., government) should make an intervention available or a third party payor (e.g., Medicare or commercial insurance) should cover an intervention. However, when the objective is to increase historically low influenza vaccination rates, accounting for only these two perspectives is myopic and may fail to account for key driving forces or stakeholders. Cost is certainly not the only (or even the primary) factor in a patient's decision to get vaccinated, but its role is salient. As direct-to-consumer marketing grows more pervasive in health care, the patient economic perspective may be increasing in importance.

A trend over the past two decades has been casting patients as “consumers,” who must choose among goods or interventions offered by traditional and non-traditional health care providers.[34] Commercials and advertisements tout the “value” of health care goods and services. While this value is sometimes quantified in terms of clinical efficacy, it is often not quantified monetarily. In many other situations where consumers have a choice (e.g., purchasing automobiles, food, or clothing), understanding the economic value of a good or service from the consumer's perspective can help both the consumers and providers of the good or service. Consumers may realize that a good is either more or less valuable than anticipated, which may in turn affect their decision-making (e.g., is it worthwhile to miss work to get vaccinated). Providers may be able to tailor the good or service to assist the consumer (e.g., making it more convenient for patients to get vaccinated or adjusting the price of the vaccine).

Although economic indicators and models cannot fully capture all the advantages and disadvantages of a medical intervention, they can offer important benchmarks, perspectives, and insights. In our model, the strongest drivers of the vaccine's economic value were the clinical attack rate followed by vaccination cost and the time required (i.e. lost productivity) by the individual to get vaccinated. This suggests that in a seasonal or a relatively mild pandemic scenario such as the 2009 H1N1 pandemic, making influenza vaccination less costly and more convenient may be an important priority and prerequisite for improved vaccination coverage.[4,35–36] For example, providing vaccination at locations the patient already frequents (e.g., work, churches, retail shops and clinics, concerts, athletic events, and other public gatherings) may be a useful cost-saving measure for individuals. Alternatively, in more severe epidemics, making a special trip and queuing for vaccination may be relatively less costly for an individual.

The relative parity in cost of vaccination observed between uninsured and insured individuals is informative as well. Finding a significant difference between the two would be expected based on reports that insurance status often dictates the decision to get vaccinated, and also would suggest that more tailored approaches to promote vaccination in each group may be needed.[4,37] Instead, the two groups seem to be facing similar costs for vaccination, i.e., lack of insurance coverage may not actually be a barrier for vaccination.

4.1 Limitations

All computer models make simplifying assumptions and cannot represent all possible outcomes that may arise from influenza vaccination or disease. Additionally, they cannot account for the vast diversity in the socio-demographic and health characteristics of the adult population. Our model only considered the healthy adult population which comprises the

majority of the population. High-risk individuals (e.g., those with major co-morbidities such as diabetes, pregnant women, or health care workers) may either have higher risk for influenza-associated morbidity and, therefore, could reap more benefits (i.e., greater cost-savings) from vaccination or could have additional reasons for being immunized (e.g., protecting neonates or patients). Future studies may focus on different categories of high-risk individuals. Although all data came from referenced sources identified from an extensive review of the literature and widely-used databases, the choice of probability distributions could bias the results. Costs drew from gamma distributions, frequently utilized for health care related costs that tend to cluster in the lower end of the distribution and have relatively fewer higher costs trail off into the upper tail. Probability parameters drew from beta distributions, frequently used to represent continuous variables that have values between 0 and 1. Lastly, parameters with relatively sparse observations drew from triangular distributions, built from a minimum, maximum, and likeliest value.[38–39] Moreover, it is difficult to quantify (both monetarily and in terms of health benefits) all facets of vaccine value, such as convenience, patient preference (for either injection or nasal spray, where indicated) and comfort, and the relative worth of LAIV and TIV for all scenarios. The decision to get the influenza vaccine and the type of vaccine administered should be determined on a case-by-case basis.

4.2 Conclusions and Future Directions

Understanding the cost-benefit of vaccination from the patient perspective can provide interesting insights for vaccination policy. While cost is certainly not the only driving factor in a patient's decision to get vaccinated, studies have suggested that a patient's perceived cost-benefit of a medical intervention could affect his or her utilization of the intervention. Aside from vaccine cost and attack rate, one of the biggest drivers of vaccination costs was a patient's time required for vaccination, so decision makers may want to focus on ways to reduce this time, such as bringing vaccination to individuals (e.g., at work, churches, or other normally frequented locations). Ideally, reducing the time demand and cost of immunization for patients will help improve influenza vaccine coverage.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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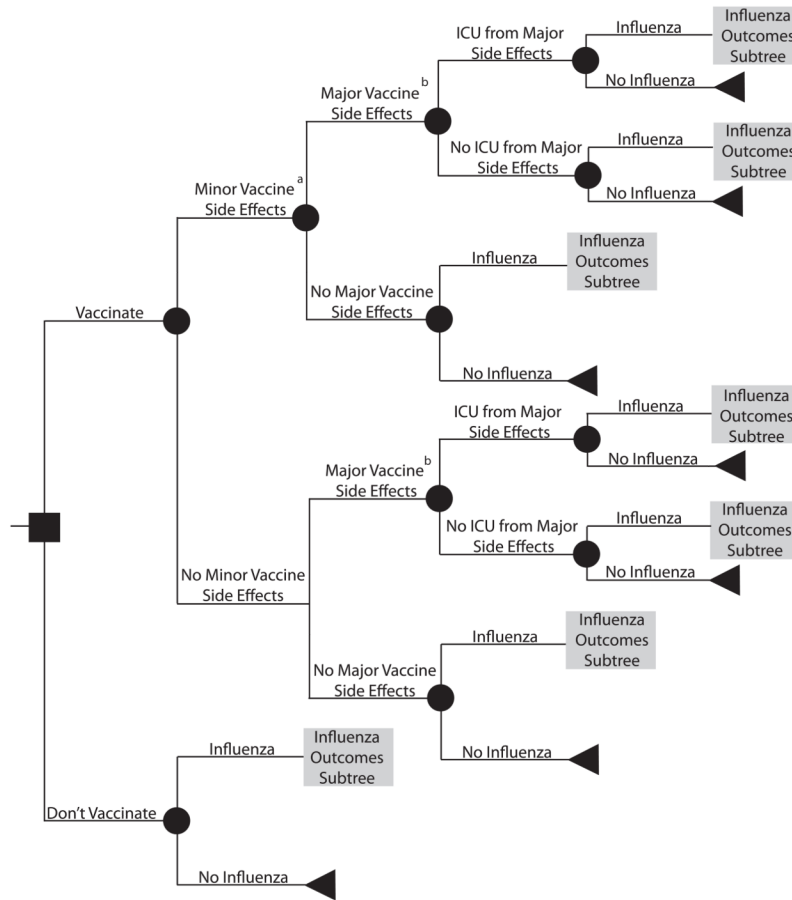


FIGURE 1. General structure of the decision model. ^a Includes local inflammation or minor systemic flu-like symptoms, requiring self-treatment with ibuprofen. ^b Guillian-Barré Syndrome (GBS)

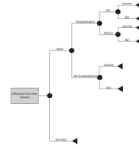


FIGURE 2.
Influenza outcomes subtree

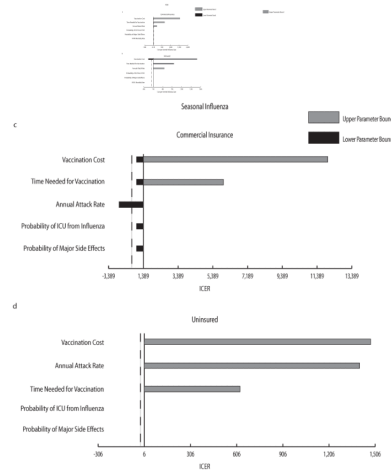


FIGURE 3.

a–d: Tornado Diagrams. a–b H1N1 vaccination scenarios for commercially insured (a) and uninsured (b) groups; c–d seasonal influenza vaccination scenarios for commercially insured (c) and uninsured (d) groups. The central axis represents the resulting incremental cost per averted influenza case when all parameters listed were set to their midpoint value of their explored ranges: vaccine cost (range: \$0 – \$150), attack rate (range: 6.5%–12.5% for H1N1 and 2%–15% for seasonal influenza), time needed for vaccination (range: 1–4 hours), major vaccine side effects (range: 0.0003%–0.0006%), ICU from influenza (15%–25% for H1N1 and 5%–20% for seasonal influenza), case fatality rate (0.011%–0.186% for H1N1). Table 1 shows baseline values for each parameter. The dashed vertical line marks where the incremental cost per averted case = 0. Negative incremental costs occurred when vaccination was economically dominant compared to no vaccination.

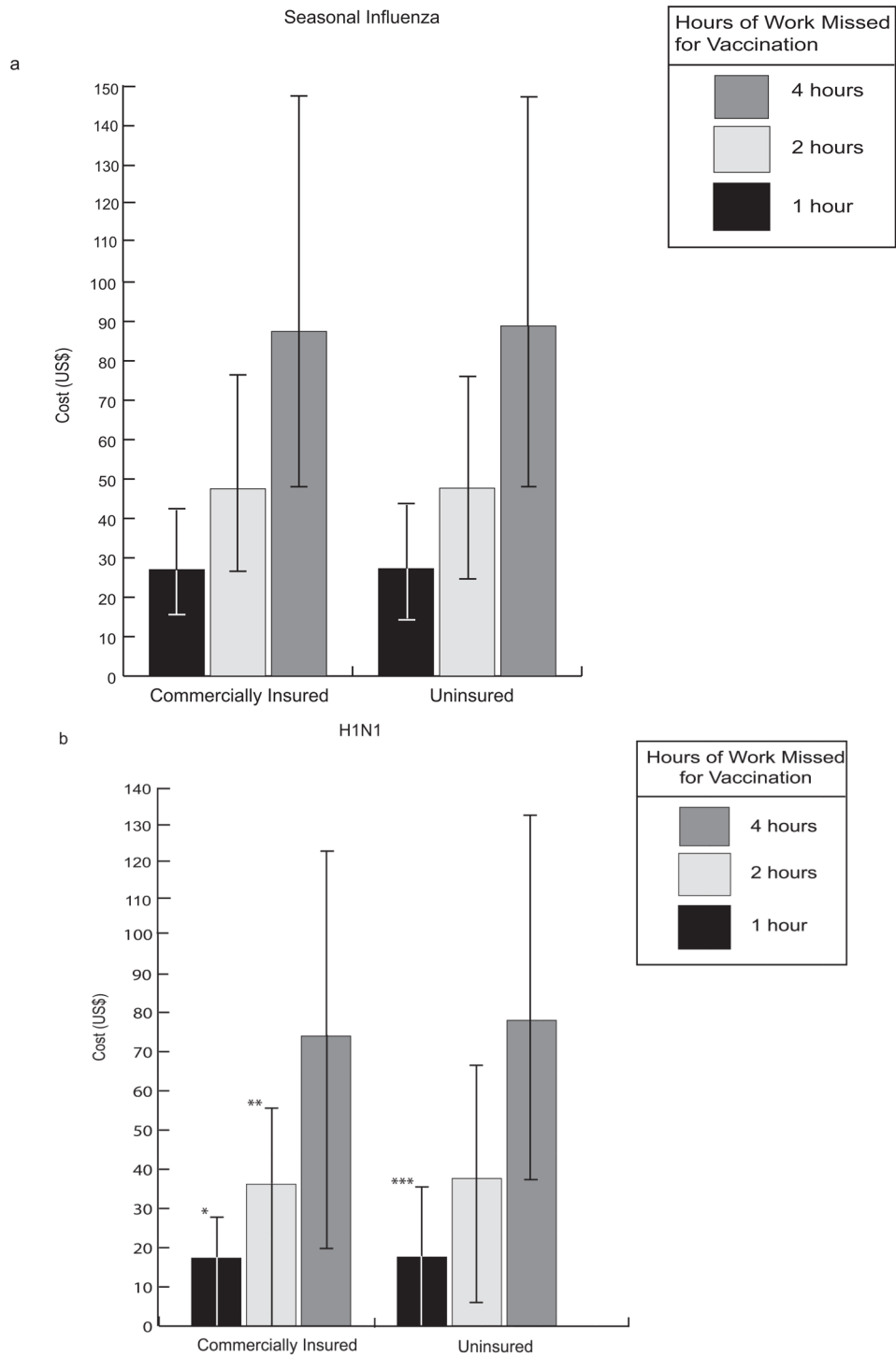


FIGURE 4. Comparison of median costs for influenza vaccination with 95% confidence interval (2.5th and 97.5th percentiles) between uninsured and commercially insured patients for (A) seasonal influenza and (B) 2009 H1N1 influenza. This data assumes a clinical attack rate of 10%, TIV cost of \$15, and baseline probabilities shown in Table 1. Negative values indicate cost-savings. * 2.5th percentile = -20.9 ** 2.5th percentile = -7.01 *** 2.5th percentile = -7.78

TABLE 1

Data Inputs for Model Variables

Variable (units)	Mean	Standard Deviation	95% Range		Distribution Type	Source
			Lower Limit	Upper Limit		
Costs (US\$)						
<i>Influenza</i>						
Home Treatment	15.61	-	11.7	19.52	Δ	[22]
Antiviral Medication (Oseltamivir)	99.32	21.99	-	-	γ	[22]
Vaccine (LAIV)	25	-	-	-	-	[22]
Vaccine TIV)	15	-	-	-	-	[22]
Clinic Cost						
Commercial Insurance	20.26	-	-	-	-	[24–25,40]
Uninsured	13.53	-	-	-	-	[24–25,40]
Hospitalization						
Commercial Insurance	1,013.41	1,195.84	-	-	γ	[24,40]
Uninsured	2,512.62	3,433.98	-	-	γ	[24–25,40]
ICU (in addition to hospitalization)						
Commercial Insurance	1,450.66	1,299.34	-	-	γ	[24,40]
Uninsured	4,721.98	3,956.70	-	-	γ	[24–25,40]
Daily Wage	128.30	-	66.08	313.04	Δ	[23]
Vaccine Side Effects						
Minor Side Effects	15.61	-	11.7	19.52	Δ	[22]
Hospitalization for Major Side Effects						

Variable (units)	Mean	Standard Deviation	95% Range		Distribution Type	Source
			Lower Limit	Upper Limit		
Commercial Insurance	1,555.01	1,529.14	-	-	γ	[24,40]
Uninsured	5,247.99	5,117.31	-	-	γ	[25,40]
ICU						
Commercial Insurance	2610	2571.57	-	-	γ	[24,40]
Uninsured	10,577.82	10,382.13	-	-	γ	[25,40]
Probabilities (%)						
Seasonal Influenza						
Influenza	6.6	1.7	-	-	β	[26]
Influenza Outcomes						
Hospitalization (given clinic visit)	0.4	-	0.1	0.7	Δ	[26,41]
Case Fatality Rate (given hospitalization)	0.9	0.3	-	--	β	[26]
Clinic Visit (given symptomatic infection)	31.3	1.4			β	[26]
ICU (given hospitalization)	15	-	-	-	-	[42]
H1N1 Influenza						
Influenza	11.5	-	6.5	19.4	Δ	[29]
Influenza Outcomes						
Hospitalization (given clinic visit)	0.56	-	-	-	-	[43]
Case Fatality Rate (given hospitalization) ^a	0.01	-	0.007	0.016	Δ	[30]
Clinic Visit (given symptomatic infection)	31.3	1.4	-	-	β	[26]
ICU (given hospitalization)	20	-	-	-	-	[44]

Variable (units)	Mean	Standard Deviation	95% Range		Distribution Type	Source
			Lower Limit	Upper Limit		
<i>Vaccines</i>						
LAIV						
Major Side Effects	0.002	-	-	-	-	[32]
Major Side Effects ICU	33	-	-	-	-	[45]
Minor Side Effects	5	-	-	-	-	[46]
Seasonal Vaccine Efficacy	62	-	45	73	Δ	[47]
H1N1 Vaccine Efficacy (LAIV)	83	-	-	-	-	[48–49]
TIV						
Major Side Effects	0.0003	-	-	-	-	[31]
Major Side Effects ICU	33	-	-	-	-	[45]
Minor Side Effects	5	-	-	-	-	[46]
Seasonal Vaccine Efficacy (TIV)	74	-	50	80	Δ	[47]
H1N1 Vaccine Efficacy (TIV)	78	-	-	-	-	[48–49]
Vaccine Reduces Hospitalization	56.5	-	45	68	Δ	[46]
Vaccine Reduces Mortality	21	-	0	42	Δ	[46]
Time (Days)						
Work Absenteeism ^b	1.03	-	0.5	1.5	Δ	[26,50–51]
Hospitalization (due to influenza)	3	-	-	-	-	[55]

^aData presents median and interval estimate

^bNumber of days absent from work due to symptomatic influenza infection

TABLE 2

Cost per influenza case avoided through vaccination with the seasonal influenza vaccine for the commercially insured and uninsured^d

Time for Vaccination (hrs)		Commercial Insurance						Uninsured					
		Attack Rate			Sensitivity Analyses			Attack Rate			Sensitivity Analyses		
		Baseline Scenario ^{b, c}		15% ^f	Baseline Scenario ^{b, c}		10% ^e	Baseline Scenario ^{b, c}		10% ^e	Baseline Scenario ^{b, c}		15% ^f
Vaccine Cost = \$0													
1 hr	LAIV	352 (-14-1,150)	1,660 (588-4,371)	161 (-85-511)	40 (-175-270)	418 (-29-1,287)	1,633 (582-4,561)	168 (-112-507)	48 (-186-299)				
	TIV	291 (-50-993)	1,400 (519-3,549)	116 (-124-410)	10 (-182-195)	126 (-40-1,033)	2,833 (1,113-7,562)	123 (-117-428)	204 (-71-535)				
2 hrs	LAIV	863 (233-2,562)	3,300 (1,350-8,529)	501 (99-1,180)	265 (-42-717)	1,018 (267-2,490)	3,409 (1,289-9,044)	511 (99-1,108)	273 (-26-684)				
	TIV	736 (198-2,139)	2,945 (1,214-7,492)	415 (61-972)	201 (-46-563)	895 (-999-2,344)	2,956 (1,229-7,310)	454 (47-989)	227 (-50-605)				
4 hrs	LAIV	1,963 (718-5,013)	6,931 (2,935-16,806)	1,229 (438-2,486)	739 (225-1,501)	2,163 (690-4,614)	6,925 (3,012-17,771)	1,207 (465-2,542)	736 (215-1,561)				
	TIV	1,669 (617-4,352)	6,120 (2,522-14,729)	1,029 (374-2,130)	599 (150-1,334)	6,768 (2,643-15,347)	6,104 (2,643-15,347)	1,006 (372-2,153)	597 (152-1,329)				
Vaccine Cost = \$15													
1 hr	TIV	617 (207-1,606)	2,505 (1,284-6,285)	332 (81-662)	156 (-51-386)	698 (210-1,614)	2,604 (1,263-6,496)	344 (65-702)	164 (-57-384)				
	TIV	1,117 (409-2,754)	4,012 (1,939-9,024)	625 (262-1,232)	348 (113-757)	1,261 (421-2,874)	4,172 (2,065-9,882)	649 (251-1,238)	363 (57-769)				
4 hrs	TIV	2,016 (787-4,830)	7,030 (3,235-16,220)	1,195 (586-2,432)	743 (307-1,512)	2,312 (841-5,104)	6,982 (3,347-17,018)	1,249 (841-5,104)	756 (307-1,451)				
Vaccine Cost = \$25													
1 hr	LAIV	1,004 (437-2,417)	3,674 (2,037-8,845)	583 (259-1,007)	315 (107-606)	1,119 (416-2,394)	3,698 (1,987-8836)	575 (262-1,016)	324 (80-606)				
	LAIV	1,545 (682-3,526)	5,545 (2,894-11,989)	911 (479-1,699)	548 (227-1,000)	1,765 (712-3,885)	5,480 (2,834-12,487)	910 (470-1,662)	554 (227-1,026)				
4 hrs	LAIV	2,696 (155-6,160)	8,886 (4,370-21,334)	1,605 (834-2,889)	1,003 (456-1,853)	2,872 (1,123-6,322)	9,002 (4,271-20,466)	1,589 (858-2991)	992 (495-1,877)				
Vaccine Cost = \$50													
1 hr	LAIV	1,653 (859-4,041)	5,964 (3,322-12,703)	1,003 (605-1,584)	594 (335-974)	1,813 (773-3,583)	5,956 (3,375-13,085)	1,001 (605-1,549)	560 (334-960)				
	TIV	1,496 (708-3,206)	5,101 (2,985-10,843)	863 (537-1,351)	497 (256-797)	1,582 (678-3,202)	5,170 (3,029-10,765)	863 (504-1,366)	506 (250-824)				
2 hrs	LAIV	2,202 (859-4,041)	7,602 (4,341-17,141)	1,332 (807-2,086)	834 (482-1,357)	2,441 (1,076-4,836)	7,618 (4,132-17,052)	1,371 (844-2,233)	840 (449-1,382)				
	TIV	1,897 (925-4,402)	6,718 (3,823-14,282)	1,153 (707-1,880)	710 (392-1,179)	2,082 (942-4,372)	6,732 (3,695-14,217)	1,161 (712-1,878)	717 (414-1,175)				
4 hrs	LAIV	3,315 (1,534-7,428)	11,601 (5,917-26,883)	1,999 (1,202-3,398)	2,180 (726-2,180)	3,639 (1,570-7,637)	11,119 (5,606-26,161)	2,026 (1,147-3,456)	1,268 (749-2,210)				
	TIV	2,946 (1,448-6,833)	9,702 (5,015-20,958)	1,769 (1,047-2,997)	1,094 (621-1,882)	3,187 (1,351-6,869)	9,941 (5,199-20,658)	1,756 (1,012-2,959)	1,125 (587-1,972)				

Time for Vaccination (hrs)		Commercial Insurance						Uninsured							
		Attack Rate						Attack Rate							
		Baseline Scenario ^{b, c}		Sensitivity Analyses		Sensitivity Analyses		Baseline Scenario ^{b, c}		Sensitivity Analyses		Sensitivity Analyses			
		2% ^d	10% ^e	15% ^f			2% ^d	10% ^e	15% ^f			2% ^d	10% ^e	15% ^f	
1 hr	LAIV	4,185 (2,371–8,709)	14,426 (9,076–32,850)	2,644 (1,899–3,933)	1,706 (1,264–2,454)	4,648 (2,371–9,300)	14,123 (8,603–30,352)	2,671 (1,941–3,845)	1,703 (1,233–2,433)						
	TIV	3,782 (2,070–8,197)	12,678 (7,815–25,186)	2,323 (1,707–3,323)	1,465 (1,101–1,465)	4,101 (2,015–7,950)	12,474 (7,624–27,624)	2,331 (1,712–3,290)	1,490 (1,098–2,084)						
2 hrs	LAIV	4,817 (2,643–10,472)	15,823 (9,451–34,041)	2,992 (2,186–4,380)	1,954 (1,386–2,810)	5,123 (2,541–9,764)	16,027 (9,206–36,616)	3,015 (2,138–4,504)	1,947 (1,422–2,842)						
	TIV	4,147 (2,211–8,919)	14,173 (5,526–29,202)	2,620 (1,850–4,028)	1,660 (1,204–2,484)	4,587 (2,295–8,884)	13,976 (8,397–29,671)	2,682 (1,885–3,852)	1,720 (1,198–2,380)						
4 hrs	LAIV	6,043 (3,196–12,655)	19,286 (11,257–42,528)	3,703 (2,501–5,683)	2,395 (1,701–3,602)	6,445 (3,125–13,004)	19,622 (11,200–42,461)	3,723 (2,573–5,660)	2,393 (1,659–3,685)						
	TIV	5,137 (2,776–11,535)	17,159 (10,200–35,421)	3,207 (2,267–4,999)	2,116 (1,466–3,182)	5,651 (10,208–36,849)	17,204 (10,208–36,849)	3,283 (2,262–5,119)	2,091 (1,453–3,095)						

^a median (95% confidence interval)

^b Beta distribution: mean = 6.6%, standard deviation = 1.7%

^c Most likely attack rate

^d Based on a distribution generated using data from Molinari et al, the probability of a 2% attack rate of seasonal influenza is < 2.5%

^e Based on a distribution created using data from Molinari et al, the probability of an influenza attack rate of 10% or greater is ≤ 2.5%

^f Based on a distribution generated using data from Molinari et al, the probability of a 15% attack rate of seasonal influenza is < 2.5%

TABLE 3

Cost per influenza case avoided through vaccination with the H1N1 vaccine for the commercially insured and uninsured^a

Time for Vaccination (hrs)	Commercial Insurance				Uninsured			
	Attack Rate		Sensitivity Analyses 6.5% ^d		Attack Rate		Sensitivity Analyses 6.5% ^d	
	Baseline Scenario ^{b, c}	6.5% ^d	Baseline Scenario ^{b, c}	6.5% ^d	Baseline Scenario ^{b, c}	6.5% ^d	Baseline Scenario ^{b, c}	6.5% ^d
Vaccine Cost = \$0								
1 hr	LAIV	16 (-187-275)	202 (-61-548)	14 (-206-263)	204 (-115-576)			
	TIV	27 (-180-295)	225 (-48-575)	29 (-228-273)	238 (-79-624)			
2 hrs	LAIV	240 (-56-690)	604 (164-1,170)	231 (-90-729)	601 (143-1,256)			
	TIV	256 (-50-728)	664 (196-1,294)	256 (-73-762)	657 (158-1,319)			
4 hrs	LAIV	675 (169-1,573)	1,407 (582-2,591)	665 (137-1,528)	1,391 (556-2,647)			
	TIV	720 (158-1,648)	1,487 (574-2,734)	723 (155-1,638)	1,502 (605-2,920)			
Vaccine Cost = \$15								
1 hr	TIV	193 (-54-508)	527 (196-919)	723 (155-1,638)	528 (168-943)			
	TIV	426 (54-986)	969 (451-1,678)	427 (64-983)	957 (413-1,682)			
4 hrs	TIV	868 (307-1,837)	1,791 (837-3,182)	865 (250-1,734)	1,792 (831-3,185)			
Vaccine Cost = \$25								
1 hr	LAIV	270 (22-654)	675 (365-1,077)	277 (5-650)	675 (318-1,100)			
	LAIV	492 (133-1,021)	1,089 (588-1,750)	484 (115-1,009)	1,069 (549-1,780)			
4 hrs	LAIV	904 (320-1,822)	1,896 (1,016-3,202)	911 (334-1,793)	1,860 (966-3,164)			
Vaccine Cost = \$50								
1 hr	LAIV	523 (212-989)	1,140 (144-1,652)	519 (195-978)	1,155 (713-1,704)			
	TIV	584 (208-1,110)	1,222 (790-1,783)	573 (223-1,127)	1,244 (777-1,824)			
2 hrs	LAIV	746 (317-1,382)	1,535 (961-2,288)	742 (316-1,434)	1,571 (1,000-2,360)			
	TIV	824 (365-1,534)	1,663 (1,051-2,558)	821 (358-1,550)	1,675 (1,038-2,534)			
4 hrs	LAIV	1,141 (519-2,128)	2,342 (1,382-3,724)	1,162 (515-2,257)	2,377 (1,339-3,874)			
	TIV	1,274 (527-2,488)	2,591 (1,526-3,908)	1,268 (572-2,440)	2,534 (1,441-4,014)			
Vaccine Cost = \$150								
1 hr	LAIV	1,543 (874-2,590)	3,027 (2,256-4,091)	1,563 (882-2,605)	3,046 (2,249-4,182)			

Time for Vaccination (hrs)	Commercial Insurance				Uninsured	
	Attack Rate		Attack Rate		Attack Rate	
	Baseline Scenario ^{b, c}	Sensitivity Analyses 6.5% ^d	Baseline Scenario ^{b, c}	Sensitivity Analyses 6.5% ^d	Baseline Scenario ^{b, c}	Sensitivity Analyses 6.5% ^d
2 hrs	TIV	1,645 (953–2,786)	3,217 (2,406–4,320)	1,657 (937–2,823)	3,265 (2,374–4,384)	
	LAI	1,769 (986–2,995)	3,430 (2,460–4,668)	1,750 (1,013–2,897)	3,425 (2,479–4,820)	
	TIV	1,837 (1,067–3,165)	3,688 (2,683–5,250)	1,870 (1,094–3,281)	3,683 (2,628–5,090)	
4 hrs	LAI	2,187 (1,221–3,901)	4,201 (2,959–5,927)	2,217 (1,182–3,921)	4,188 (2,903–5,968)	
	TIV	2,356 (1,302–4,174)	4,533 (3,109–6,322)	2,387 (1,305–4,201)	4,481 (3,049–6,368)	

^a median (95% confidence interval)

^b Triangular distribution: Min = 6.5%, Likeliest = 11.5%, Max = 19.4

^c Most likely attack rate

^d Based on a distribution generated from data collected by Miller et al, the probability of a 6.5% attack rate of seasonal influenza is approximately <1