

Ranking Vaccines

A Prioritization Framework

Phase I: Demonstration of Concept
and a Software Blueprint



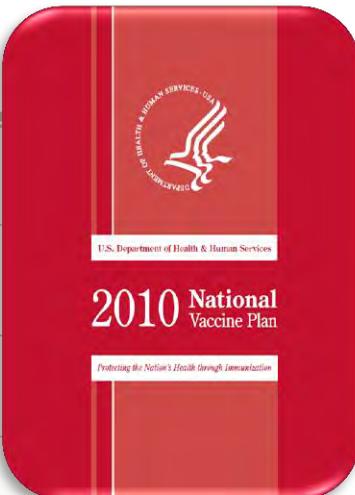
INSTITUTE OF MEDICINE

OF THE NATIONAL ACADEMIES

Advising the nation • Improving health

National Vaccine Plan Priorities for Implementation

- A. Develop a catalogue of priority vaccine targets of domestic and global health importance (Goal 1).
- B. Strengthen the science base for the development and licensure of new vaccines (Goals 1 and 2).
- C. Enhance timely detection and verification of vaccine safety signals and develop a vaccine safety scientific agenda (Goal 2).
- D. Increase awareness of vaccines, vaccine-preventable diseases, and the benefits/risks of immunization among the public, providers, and other stakeholders (Goal 3).



A.

Develop a catalogue of priority vaccine targets of domestic and global health importance (Goal 1).

access to routinely recommended vaccines (Goal 4).

- G. Create an adequate and stable supply of routinely recommended vaccines and vaccines for public health preparedness (Goal 4).
- H. Increase and improve the use of interoperable health information technology and electronic health records (Goal 4).
- I. Improve global surveillance for vaccine-preventable diseases and strengthen global health information systems to monitor vaccine coverage, effectiveness, and safety (Goal 5).
- J. Support global introduction and availability of new and under-utilized vaccines to prevent diseases of public health importance (Goal 5).

Vision of the National Vaccine Program Office

Step 1

Create and Validate the Model

Phase I

Enhance the Model

Phase II

Step 2

Execute and Populate the Model

Step 3

Prioritize Vaccines and Evaluate the Catalogue Against Ideal Attributes

Charge to the Committee

1. Review domestic and global research and development prioritization activities relevant to identifying new preventive vaccine targets.
2. Develop an analytical framework and model for prioritizing vaccines of domestic and global importance. Engage stakeholders to inform the process of the model development and implementation.
3. Test and validate the model using two to three predetermined vaccines, including at least one vaccine candidate of domestic importance and one of global importance.
4. Prepare a report containing the analytical framework and model for evaluating and prioritizing vaccine targets along with recommendations as to how to use the model for reviewing the catalog of preventive vaccines every two to three years.

Previous IOM Reports

New Vaccine Development
Establishing Priorities

VOLUME I

Diseases of Importance in the United States



NAS
RRAE
IOM

1985

New Vaccine Development
Establishing Priorities

VOLUME II

Diseases of Importance in
Developing Countries



NAS
RRAE
IOM

1986

Vaccines
for the 21st Century

A TOOL FOR DECISIONMAKING

INSTITUTE OF MEDICINE

2000

The Committee Approach

16

Expert
Members

5

Committee
Meetings

2

Public
Meetings

n

Subgroup
Meetings

2

2

11

3

Modeling
Consultants

Software
Designers

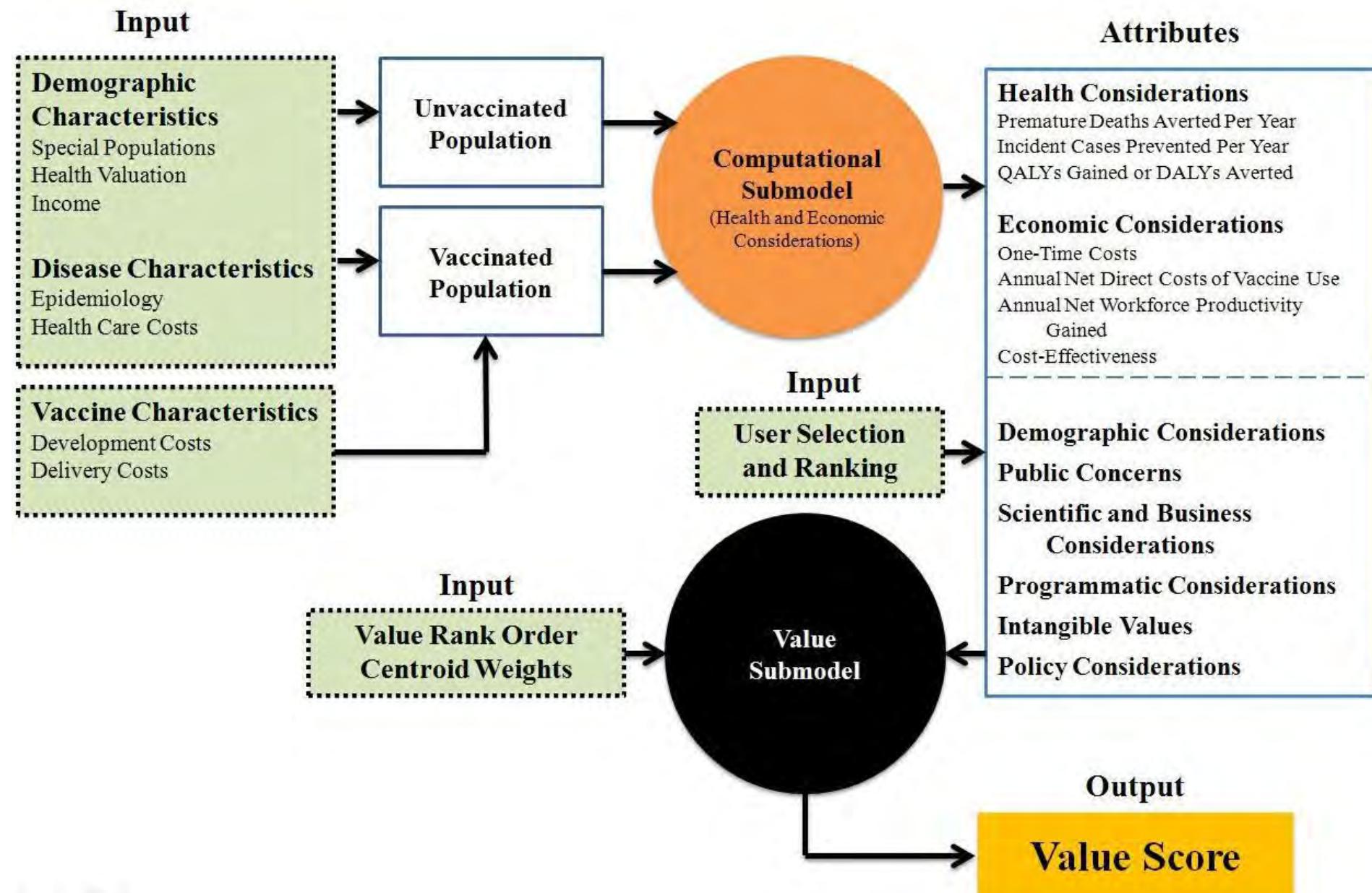
Concept
Evaluators

Research
Staff
Members

Modeling Considerations

1. Multi-Stakeholder Application
2. Axiomatic Foundation
3. Relative Priority Scaling
4. Sensitivity Analysis
5. Transparency
6. User-Friendly Software Base

The Multi-Attribute Framework



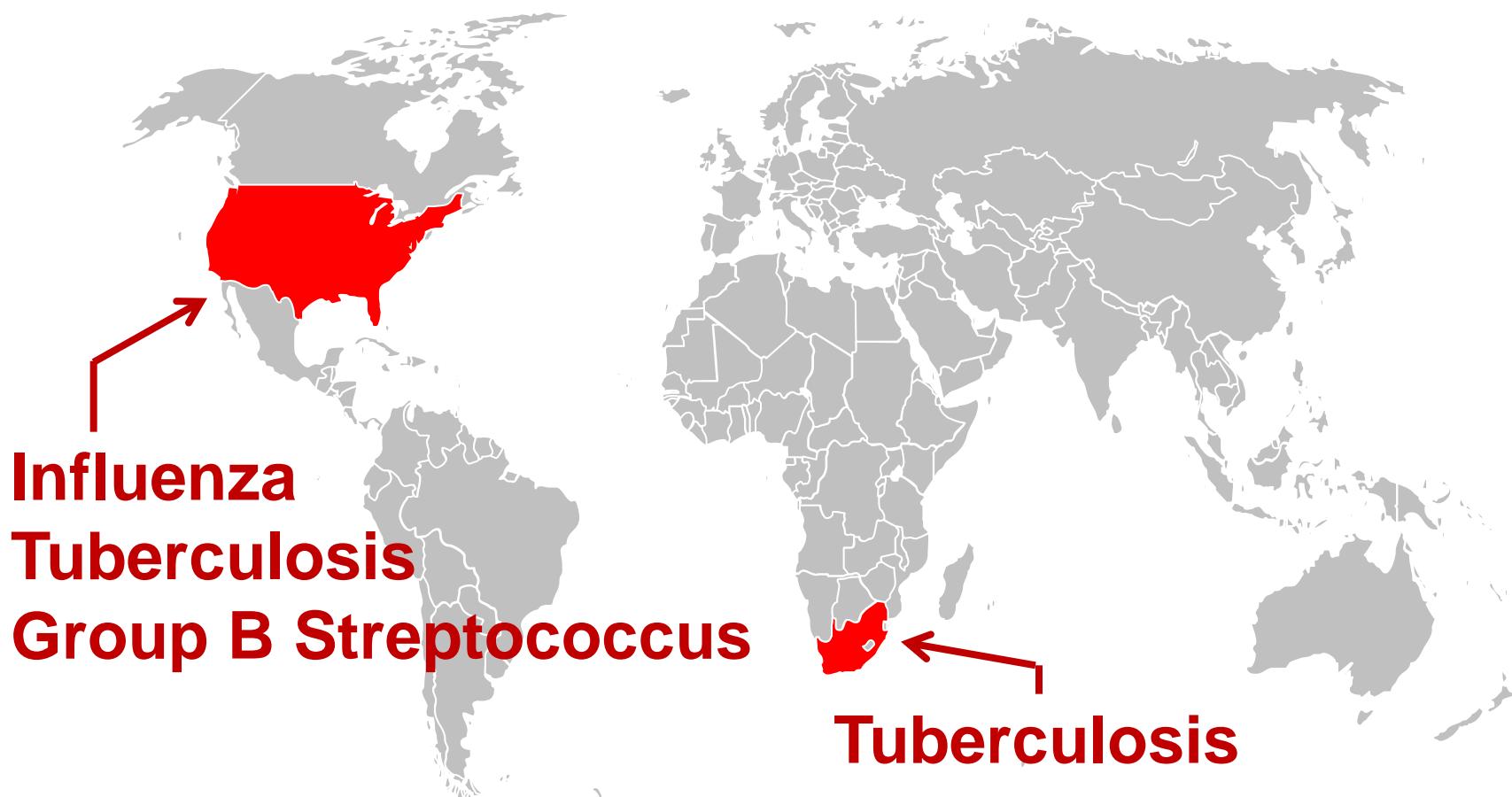
Attribute Selection

8

29

Health Considerations	<ul style="list-style-type: none">• Premature Deaths Averted Per Year• Incident Cases Prevented Per Year• QALYs Gained or DALYs Averted
Economic Considerations	<ul style="list-style-type: none">• One-Time Costs• Annual Net Direct Costs (Savings) of Vaccine Use• Annual Net Workforce Productivity Gained• Cost-Effectiveness
Demographic Considerations	<ul style="list-style-type: none">• Benefits Infants and Children• Benefits Women• Benefits Socioeconomically Disadvantaged• Benefits Military Personnel• Benefits Other Priority Population
Public Concerns	<ul style="list-style-type: none">• Availability of Alternative Public Health Measures• Potential Complications Due to Vaccines• Disease Raises Fear and Stigma in the Public• Serious Pandemic Potential
Scientific and Business Considerations	<ul style="list-style-type: none">• Likelihood of Financial Profitability for the Manufacturer• Likelihood of Successful Licensure in 10 Years• Demonstrates New Production Platforms• Existing or Adaptable Manufacturing Techniques• Potential Litigation Barriers Beyond Usual• Interests from NGOs and Philanthropic Organizations
Programmatic Considerations	<ul style="list-style-type: none">• Potential to Improve Delivery Methods• Fits into Existing Immunization Schedules• Reduces Challenges Relating to Cold-Chain Requirements
Intangible Values	<ul style="list-style-type: none">• Eradication or Elimination of the Disease• Vaccine Raises Public Health Awareness
Policy Considerations	<ul style="list-style-type: none">• Special Interest for National Security, Preparedness, and Response• Advances Nation's Foreign Policy Goals

New Vaccine Candidates



Rank Order Centroid Weighting

Rank	Number of Attributes Selected													
	2	3	4	5	6	7	8	9	10	11	12	13	14	15
1*	0.750	0.611	0.521	0.457	0.408	0.370	0.340	0.314	0.293	0.275	0.259	0.245	0.232	0.221
2	0.250	0.278	0.271	0.257	0.242	0.228	0.215	0.203	0.193	0.184	0.175	0.168	0.161	0.155
3		0.111	0.146	0.157	0.158	0.156	0.152	0.148	0.143	0.138	0.134	0.129	0.125	0.121
4			0.063	0.090	0.103	0.109	0.111	0.111	0.110	0.108	0.106	0.104	0.101	0.099
5				0.040	0.061	0.073	0.079	0.083	0.085	0.085	0.085	0.084	0.083	0.082
6					0.028	0.044	0.054	0.061	0.065	0.067	0.068	0.069	0.069	0.069
7						0.020	0.033	0.042	0.048	0.052	0.054	0.056	0.057	0.058
8							0.016	0.026	0.034	0.039	0.043	0.045	0.047	0.048
9								0.012	0.021	0.027	0.032	0.036	0.038	0.040
10									0.010	0.017	0.023	0.027	0.030	0.033
11										0.008	0.015	0.019	0.023	0.026
12											0.007	0.012	0.017	0.020
13												0.006	0.011	0.014
14													0.005	0.009
15														0.004
Total**	1.00	1.00	1.00	1.00	1.00	1.00	1.00	1.00	1.00	1.00	1.00	1.00	1.00	1.00

$$w_i = \sum_{j=i}^n \frac{1}{j} \quad i = 1 \dots n$$

*Highest rank = 1

**Totals may not add to 1.00 due to rounding

SMART Vaccines Beta

Step 0 of 7 Terms and Conditions

Disclaimer

SMART Vaccines Beta

A Prototype Framework for Prioritizing New Vaccines

Phase I: Demonstration of Concept

April 2012

Committee on Identifying and Prioritizing New Preventive Vaccines for Development
Institute of Medicine

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The Institute of Medicine (IOM) Committee on Identifying and Prioritizing New Preventive Vaccines for Development (Committee) of the National Academy of Sciences (NAS) is tasked with developing an analytical framework and model for prioritizing vaccines of domestic and global importance, and to engage stakeholders to inform the process of the model development and implementation. The Committee, with the assistance of consultants from Johns Hopkins University and VIM Interactive, has developed, as part of Phase I of the study, a prototype software entitled "SMART Vaccines Beta" which is ultimately intended to be a decision-assist tool and not a decision maker. In its current version, this prototype is NOT usable to assist any decision-making process. Subsequent work will be focused on improving the prototype software.

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New
Users

Returning
Users

To proceed, enter the following:

Your Name

Your Institution

Your Email Address

password

I agree to the terms and conditions of use above.

**Proceed to SMART
Vaccines Beta**

Disclaimer

SMART Vaccines is intended to serve only as a **decision-support tool** and should *not* be used as a decision maker.

SMART Vaccines Beta

Step 1 of 7
Selection and
Ranking of Values

Values

Demographics

Disease Burden

Vaccines

Value Assessment

Value Score

Step 1: Select Vaccine Values and Rank Their Importance

[To Demographics](#)

HEALTH CONSIDERATIONS

- Premature Deaths Averted Per Year
- Incident Cases Prevented Per Year
- QALYs Gained or DALYs Averted Per Year

ECONOMIC CONSIDERATIONS

- One-Time Costs
- Annual Net Direct Costs of Vaccine Use
- Annual Net Workforce Productivity Gained
- Cost Effectiveness

POLICY CONSIDERATIONS

- Special Interest for National Security, Preparedness, and Response
- Potential Complications Due to Vaccines

PUBLIC CONCERN

- Availability of Alternative Public Health Measures
- Potential Complications Due to Vaccines
- Disease Raises Fear and Stigma in the Public
- Serious Pandemic Potential

DEMOGRAPHIC CONSIDERATIONS

- Benefits Infants and Children
- Benefits Women
- Benefits Socioeconomically Disadvantaged
- Benefits Military Personnel
- Benefits Other Priority Population

PROGRAMMATIC CONSIDERATIONS

- Potential to Improve Delivery Methods
- Fits into Existing Immunization Schedules
- Reduces Challenges Relating to Cold-Chain Requirements

SCIENTIFIC AND BUSINESS CONSIDERATIONS

- Likelihood of Financially Profitability for the Manufacturer
- Likelihood of Successful Licensure in 10 Years
- Demonstrates New Production Platforms
- Existing or Adaptable Manufacturing Techniques
- Potential Litigation Barriers Beyond Usual
- Interests from NGOs and Philanthropic Organizations

INTANGIBLE VALUES

- Eradication or Elimination of the Disease
- Vaccine Raises Public Health Awareness

Drop values here

(Drag to rank in order of importance)

THE NATIONAL ACADEMIES

Advisers to the Nation on Science, Engineering, and Medicine

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SMART Vaccines Beta

Step 2 of 7
Population Profile

Step 2: Enter Population Data

[Back](#)[To Disease Burden](#)

Add Population

Edit Existing Population:
- Select Population -

Save as new...

Untitled Profile

United States (US)

- Select Life Table -
Blank Template
United States (US)
South Africa (ZA)

Female Male Special

MALE

LIFE TABLE

AGE	POPULATION	Living (lx)	Life-Years (nLx)	Life Expectancy (ex)	Standard Life Expectancy (sx)	HRQoL (HUI2)	PRODUCTIVITY
< 1	2,294,679	100,000	99,348	76.0	76.0	0.99	\$ 17.90
1-4	8,889,066	99,276	396,817	75.6	75.6	0.99	\$ 17.90
5-9	10,753,934	99,156	495,604	71.7	71.7	0.99	\$ 17.90
10-14	10,838,788	99,085	495,185	66.7	66.7	0.99	\$ 17.90
15-19	11,472,812	98,989	493,905	61.8	61.8	0.99	\$ 16.80
20-24	11,374,397	98,573	491,150	57.0	57.0	0.89	\$ 16.80
25-29	11,021,998	97,887	487,775	52.4	52.4	0.89	\$ 16.80
30-34	10,581,472	97,223	484,373	47.7	47.7	0.89	\$ 16.80
35-39	10,547,351	96,526	480,477	43.1	43.1	0.89	\$ 15.49
40-44	10,872,790	95,665	475,151	38.4	38.4	0.89	\$ 15.49
45-49	11,447,885	94,396	467,208	33.9	33.9	0.84	\$ 15.49
50-54	10,825,136	92,487	455,327	29.6	29.6	0.84	\$ 15.49
55-59							

SMART Vaccines Beta

Step 3 of 7
Disease Profile



Values

Demographics

Disease Burden

Vaccines

Value Assessment

Value Score

Step 3: Enter Disease Burden and Costs Data

[Back](#)[To Vaccines](#)[Add Disease](#)

Edit Disease:

[- Select Disease -](#)[- Select Disease -](#)

- Influenza
- Tuberculosis
- Group B Strep

Influenza

[Health](#)[Economic](#)[Female](#)[Male](#)[Special](#)

Age	Population Size	Annual Incidence (per 100,000)	Case Fatality Rate (%)	Herd Immunity Threshold (%)
< 1	4,478,198	20,300	0.004 %	100 %
1 – 19	81,859,350	10,200	0.001 %	100 %
20 – 64	188,118,413	6,600	0.072 %	100 %
> 65	40,093,919	9,000	1.17 %	100 %



Values

Demographics

Disease Burden

Vaccines

Value Assessment

Value Score

Step 3: Enter Disease Burden and Costs Data

[Back](#)[To Vaccines](#)[Add Disease](#)

Edit Disease:

[Save as new...](#)[Save](#)

Influenza

[Health](#)[Economic](#)

Morbidity	Cases (%)	Disutility (toll)	Disability Weight	Duration (days)
Influenza illness without outpatient visit (D1)	59.5 %	0.09	0.01	4
Influenza illness with outpatient visit (D2)	40 %	0.13	0.1	4
Influenza hospitalization (D3)	0.5 %	0.2	0.3	4

[Add Morbidity](#)

Permanent Impairment	Cases (%)	HRQoL (HUI2)	Disability Weight
Impairment 1 (P1)	0 %	0	0
Untitled Impairment	0 %	0	0

[Add Impairment](#)

Health Care	Cost per unit	Death units	D1 units	D2 units	D3 units
Over-the-Counter medications	\$ 3.00	1	1	1	1
Physician visit	\$ 100.00	0	0	0	0



Values

Demographics

Disease Burden

Vaccines

Value Assessment

Value Score

Step 3: Enter Disease Burden and Costs Data

[Back](#)[To Vaccines](#)[Add Disease](#)

Edit Disease:

[Save as new...](#)[Save](#)

Influenza

[Health](#)[Economic](#)

Influenza illness with outpatient visit (D2)

40	%	0.13	0.1	4
----	---	------	-----	---

Influenza hospitalization (D3)

0.5	%	0.2	0.3	4
-----	---	-----	-----	---

Add Morbidity

Health Care	Cost per unit	Death units	D1 units	D2 units	D3 units
Over-the-Counter medications	\$ 3.00	1	1	1	1
Physician visit	\$ 100.00	0	0	0	0
Outpatient visit	\$ 250.00	1	0	1	1
Emergency department visit	\$ 750.00	0	0	0	0
Hospitalization	\$ 1,200.00	5	0	0	5
Total		\$6,253.00	\$3.00	\$253.00	\$6,253.00

Add Service

SMART Vaccines Beta

Step 4 of 7
Vaccines Profile

Step 4: Enter Vaccine-Specific Data

[Back](#)[To Value Assessment](#)

Load Existing Disease:

- Select Disease -

Influenza

Vaccines [+ Add](#)

Untitled Vaccine 1

Age	Total Population	Target Population (%)	Vaccine Coverage (%)	Vaccine Effectiveness (%)
< 1	4,478,197	2,283,880	40 %	60 %
1 – 19	81,859,350	41,748,269	40 %	70 %
20 – 64	188,118,413	95,940,391	40 %	75 %
> 65	40,093,919	20,447,899	40 %	40 %

[Save as new...](#) [Save](#)



Step 4: Enter Vaccine-Specific Data

[Back](#)[To Value Assessment](#)

Load Existing Disease:

- Select Disease -

Influenza

Vaccines [+ Add](#)

Untitled Vaccine 1

[Save as new...](#) [Save](#)

Vaccine Characteristics	
Length of Immunity	1 years of life
Doses Required Per Person	1 doses
Cost Per Dose	\$ 10
Cost to Administer Per Dose	\$ 15
Research Costs (approximate)	\$ 100,000,000
Licensure Costs (approximate)	\$ 500,000,000
Start-up Costs (approximate)	\$ 100,000
Time to adoption	5 years

Step 4: Enter Vaccine-Specific Data

[Back](#)[To Value Assessment](#)

Load Existing Disease:

- Select Disease -

Influenza

Vaccines

[+ Add](#)

✖ Untitled Vaccine 1

Population

Product Profile

Complications

Morbidity	Cases (%)	Disutility (toll)	Disability Weight	Duration (days)
Guillain-Barré Syndrome (A1)	0.000001 %	0.35	0.44	4
Systemic reaction (fever or achiness) (A2)	0.011 %	0.25	0.1	4
Anaphylaxis (A3)	0.00000025 %	0.25	0.44	4

Add Morbidity

Permanent Impairment	Cases (%)	HRQoL (HUI2)	Disability Weight
None	0 %	0	0

Add Impairment

Deaths	Cases (%)
None	0 %

[Save as new...](#)[Save](#)



Step 4: Enter Vaccine-Specific Data

[Back](#)[To Value Assessment](#)

Load Existing Disease:

- Select Disease -

Influenza

Vaccines [+ Add](#)

Untitled Vaccine 1

[Save as new...](#) [Save](#)

Population Product Profile Complications

Health Care	Cost per unit	A1	A2	A3
Over-the-Counter medications	\$ 3.00	0	0	0
Physician visit	\$ 100.00	0	1	0
Outpatient visit	\$ 250.00	0	0	0
Emergency department visit	\$ 750.00	0	0	1
Hospitalization	\$ 1,200.00	40	0	0
Total		\$48,000	\$100	\$750

Add Service

SMART Vaccines Beta

Step 5 of 7

Qualitative Attributes



Step 5: Assess Values

[Back](#)[To Value Score](#)

Load Existing Disease:

- Select Disease -

Influenza

Vaccine A

Vaccine B

Vaccine C

Vaccine D

Vaccine E

Vaccine F

Save as new...

Health Considerations		
Values	Assessments	
	International	U.S.A.
Premature Deaths Averted Per Year	<input checked="" type="radio"/> > 1,000,000 <input type="radio"/> 500,000 – 999,999 <input type="radio"/> 100,000 – 499,999 <input type="radio"/> < 100,000	<input checked="" type="radio"/> > 20,000 <input type="radio"/> 5,000 – 19,999 <input type="radio"/> 1,000 – 4,999 <input type="radio"/> < 1,000
Incident Cases Prevented Per Year	<input checked="" type="radio"/> > 10 million <input type="radio"/> 1 – 10 million <input type="radio"/> 250,000 – 1 million <input type="radio"/> < 250,000	The difference in the number of incident cases of disease in one year assuming no routine vaccine use and assuming routine vaccine use against the disease in the population.
Economic Considerations	<input type="button" value="+"/>	
Demographic Considerations	<input type="button" value="+"/>	
Public Concerns	<input type="button" value="+"/>	
Scientific and Business Considerations	<input type="button" value="+"/>	
Programmatic Considerations	<input type="button" value="+"/>	
Intangible Values	<input type="button" value="+"/>	
Policy Considerations	<input type="button" value="+"/>	



Step 5: Assess Values

[Back](#)[To Value Score](#)

Load Existing Disease:
- Select Disease -

Influenza

Vaccine A

Vaccine B

Vaccine C

Vaccine D

Vaccine E

Vaccine F

Sum of development plus licensure plus start-up costs. This attribute represents the magnitude of financial barriers to bringing the vaccine to use in the population.

	Assessments	
	International	U.S.A.
One-Time Costs	<input checked="" type="radio"/> < \$100 million	<input type="radio"/> < \$100 million
	<input type="radio"/> \$100 – \$500 million	<input type="radio"/> \$100 – \$500 million
	<input type="radio"/> \$500 million – \$1 billion	<input checked="" type="radio"/> \$500 million – \$1 billion
	<input type="radio"/> > \$1 billion	<input type="radio"/> > \$1 billion
Annual Net Direct Costs Savings of Vaccine Use	<input type="radio"/> >10 million	<input type="radio"/> >10 million
	<input checked="" type="radio"/> 1 – 10 million	<input checked="" type="radio"/> 1 – 10 million
	<input type="radio"/> 250,000 – 1 million	<input type="radio"/> 250,000 – 1 million
	<input type="radio"/> < 250,000	<input type="radio"/> < 250,000

Demographic Considerations

Public Concerns

Scientific and Business Considerations

Programmatic Considerations

Intangible Values

Policy Considerations

[Save as new...](#) [Save](#)

SMART Vaccines Beta

Step 6 of 7
Computed Results

Step 6: Compare Your Candidate Vaccines

[Back](#)[Rank Vaccine Values](#)

Model Unit Values

Discounting %

DALY Age Weight

Vaccine List

Check two or more vaccines to compare.

Influenza Vaccine A

Tuberculosis Vaccine B

Group B Strep Vaccine C

	Vaccine A	Vaccine B	Vaccine C
Health Considerations			
Premature Deaths Averted Per Year	100	100	100
Incident Cases Prevented Per Year	101,833,848	101,833,848	101,833,848
QALYs Gained or DALYS Averted	2,352,850	2,352,850	2,352,850
Economic Considerations			
One-Time Costs	\$600,000,000	\$600,000,000	\$600,000,000
Annual Net Direct Costs of Vaccine Use	\$100,000,000	\$100,000,000	\$100,000,000
Annual Net Workforce Productivity Gained	\$500,000,000	\$500,000,000	\$500,000,000
Cost Effectiveness	\$100,000	\$100,000	\$100,000
Demographic Considerations			
Benefits Infants and Children	Yes	Yes	Yes
Benefits Women	Yes	Yes	Yes
Benefits Socioeconomically Disadvantaged	Yes	Yes	Yes
Benefits Military Personnel	Yes	Yes	Yes
Benefits Other Priority Population	No	No	No

SMART Vaccines Beta

Step 7 of 7
Priority Score

Value Score: Results of Your Comparison

[Back](#)

Candidate Vaccines	Disease	Value Score
Vaccine A	Influenza	51.1
Vaccine B	Tuberculosis	27.8
Vaccine C	Group B Streptococcus	11.1



Drag Vaccine Values to Rank and Update Value Score

Premature Deaths Averted Per Year

One-Time Costs

Vaccine Raises Public Health Awareness

Benefits Infants and Children

Demonstrates New Production Platforms



Serious Pandemic Potential

Sample Scenarios

- Same attributes but different *values*
 - A Health Minister vs. PAHO
 - NIH vs. A Vaccine Manufacturer
- Same values but different *attributes*
 - One dose vs. Three dose Vaccines
 - Risk of Development vs. Cold-Chain Requirements

Additional Considerations

Data Demands and Quality

Subjective Weights (approximated)

Distribution Around Data

Steady-State Population

Disease Interactions (e.g., HIV+TB)

Phase II: SMART Vaccines 1.0

Public Feedback (Workshop)

Model Refinement

Software Enhancement

Additional Vaccine Candidates

Sensitivity and Uncertainty

Usability Evaluation

Recommendations

Ranking Vaccines

A Prioritization Framework



Phase I: Demonstration of Concept and a Software Blueprint

Report PDF
available at

www.nap.edu

Feedback
SMARTVaccines@nas.edu

Thank you