

# AN EVALUATION OF THE RISK DECISION LEVEL FOR PROPHYLAXIS AND TREATMENT AFTER AN ANTHRAX RELEASE



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# Acknowledgements

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# OBJECTIVES

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To link the technical research on bio-threats conducted in other CAMRA projects with the societal goal of managing the risk of bioterrorism

## Current Research Questions

1. What is the decision level for anthrax risk?
2. What dose corresponds to this risk level?  
How do we use animal dose-response studies to inform this estimate?
3. How do we relate what we measure in the environment to dose and risk?

# Flow of Information

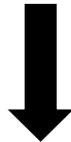
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Decision model - risk at which action is warranted



*Bacillus anthracis*  
Dose-Response Model – dose producing risk at which action is warranted

*Bacillus anthracis* Aerosol  
Transport Model-environmental concentrations producing dose which produces risk at which action is warranted



Minimum Sampling Area/Volume  
Such that a negative sample establishes that environmental concentrations would not produce a dose which produces a risk at which action is warranted



Response Decision

# Decision level for anthrax risk

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- ❑ Evaluate the cost-effectiveness of strategies for prophylaxis and treatment after an aerosolized release of *B. anthracis* (Fowler et al. 2005)
- ❑ Risk of infection is variable based on the size of the release and/or the amount of exposure to which a person is subjected
- ❑ In many situations there will be a few highly exposed individuals and a much larger number of individuals who receive much lower exposures
- ❑ At what point is medical treatment not justified?

Fowler, RA, Sanders, GD, Bravata, DM, Nouri, B, et al. 2005. Cost-Effectiveness of Defending against Bioterrorism: A Comparison of Vaccination and Antibiotic Prophylaxis against Anthrax, *Annals of Internal Medicine*, 142:601-610

# Exposure Scenarios

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- Prospective Scenario
  - ▣ Post-event re-occupancy of a building
  - ▣ Spores have settled and deposited on tracked surfaces
- Retrospective Scenario
  - ▣ *Bacillus anthracis* spores have been released in an indoor venue
  - ▣ People in the immediate vicinity will receive treatment
  - ▣ People on the outskirts may or may not require treatment (in other rooms or outside of the building)

# Methods

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- Decision analytic model (Fowler et al. 2005)
  - Societal perspective for costs and benefits
  - Discounted at 3% annually
  - Monetize remaining expected lifespan of an individual
  - All costs are in 2004 dollars
- Precision Tree 1.0 for Excel
  - Expected Value of Costs and Utilities

# Model Inputs

•Probabilities	
• <i>Clinical inhalational anthrax after attack and exposure:</i>	
•No vaccination, no antibiotics	0.95
•No vaccination, receive antibiotics	0.2
•Vaccination, no antibiotics	0.07
•Vaccination, receive antibiotics	0.02
•Baseline mortality given clinical disease	0.45
•Nondisabled state if survive clinical illness	0.85

<b>Costs, 2004 \$</b>	<b>\$</b>
Vaccine costs (6 doses)	18
Vaccine administration	46
<i>Antibiotic and administration costs (adult dosing):</i>	
Doxycycline, 100 mg, orally twice daily	12
Severe inhalational anthrax estimated cost of care	28,731
Death from any cause	6,270

# Model Inputs

<b>Utilities</b>	
Population baseline	0.92
Antibiotic treatment	0.90
Severe inhalational anthrax	0.64
Post anthrax healthy state	0.9
Post anthrax disabled state	0.8
Vaccine & Antibiotic Side Effects:	
Mild	0.9
Moderate	0.8
Severe	0.6

## **Baseline Case Assumptions for Hypothetical Cohort**

- Reside or work in metropolitan U.S. area like New York City
- Mean age = 36 years
- Life expectancy = 76 years
- Value of a QALY = \$50,000
- Utilized least expensive medication
- Anthrax related illness is severe

## Summary of Base Case Utilities

# Valuing Side Effects

- FOR ANTIBIOTICS –Reduced utilities were considered for a period of 60 days for mild and moderate and 7 days for severe side effects
- FOR VACCINATION –Reduced utilities were considered for a period of 7 days for mild and moderate and 21 days for severe side effects

# Valuing Side Effects

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**AN EXAMPLE CALCULATION for a person who receives vaccination, does not get anthrax related illness, but suffers severe side effects**

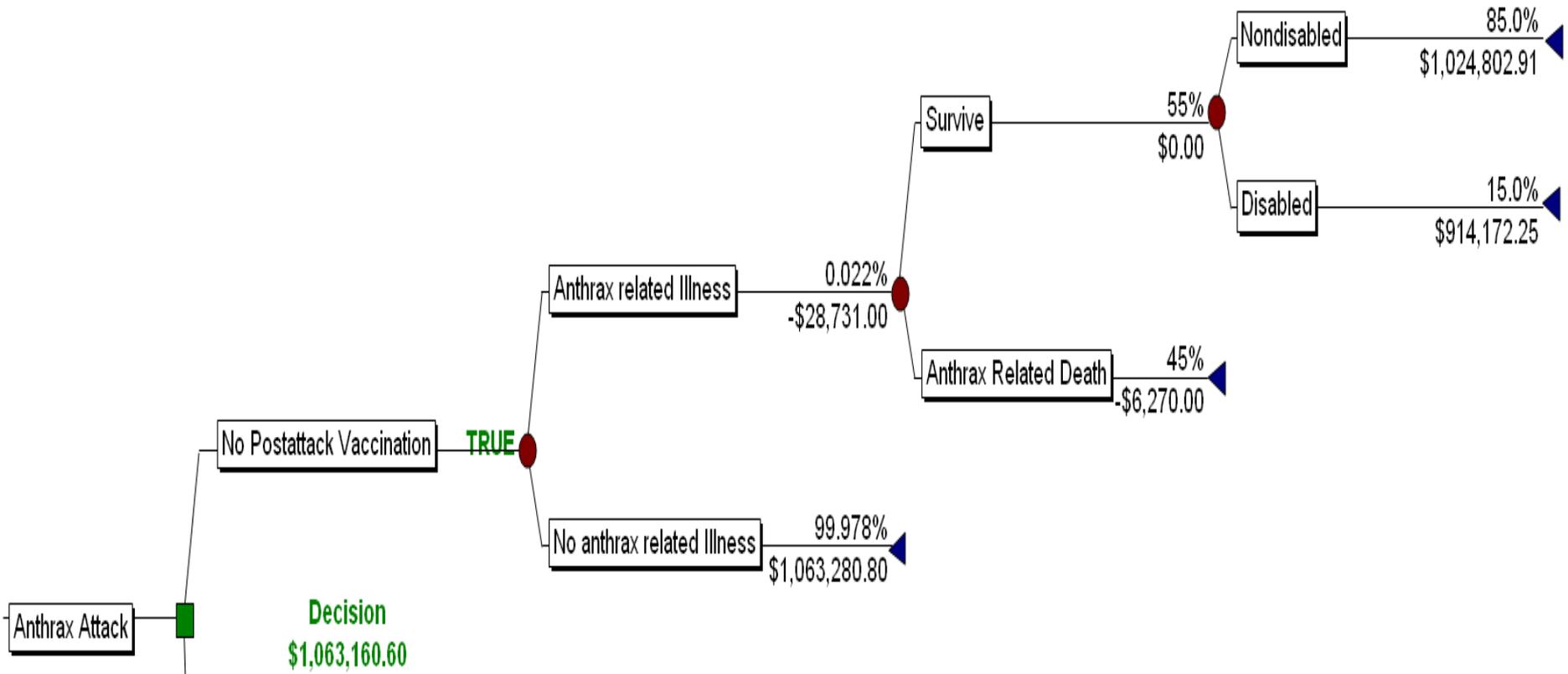
Expected Value =

- Present value of the Cost of Treatment
- + Present Value of the (Monetized QALY at the Population Baseline Utility for 39 years )
- + (Monetized QALY at Reduced Utility for 21 days + Monetized QALY at the Population Baseline Utility for the Remaining Days of the Year)

$$EV = -2473 + (((0.92 * 22.7893 * \$50,000) + ((0.6) * \$50,000 (21/365)) + (0.92 * \$50,000 * (344/365)))) * 0.9709$$

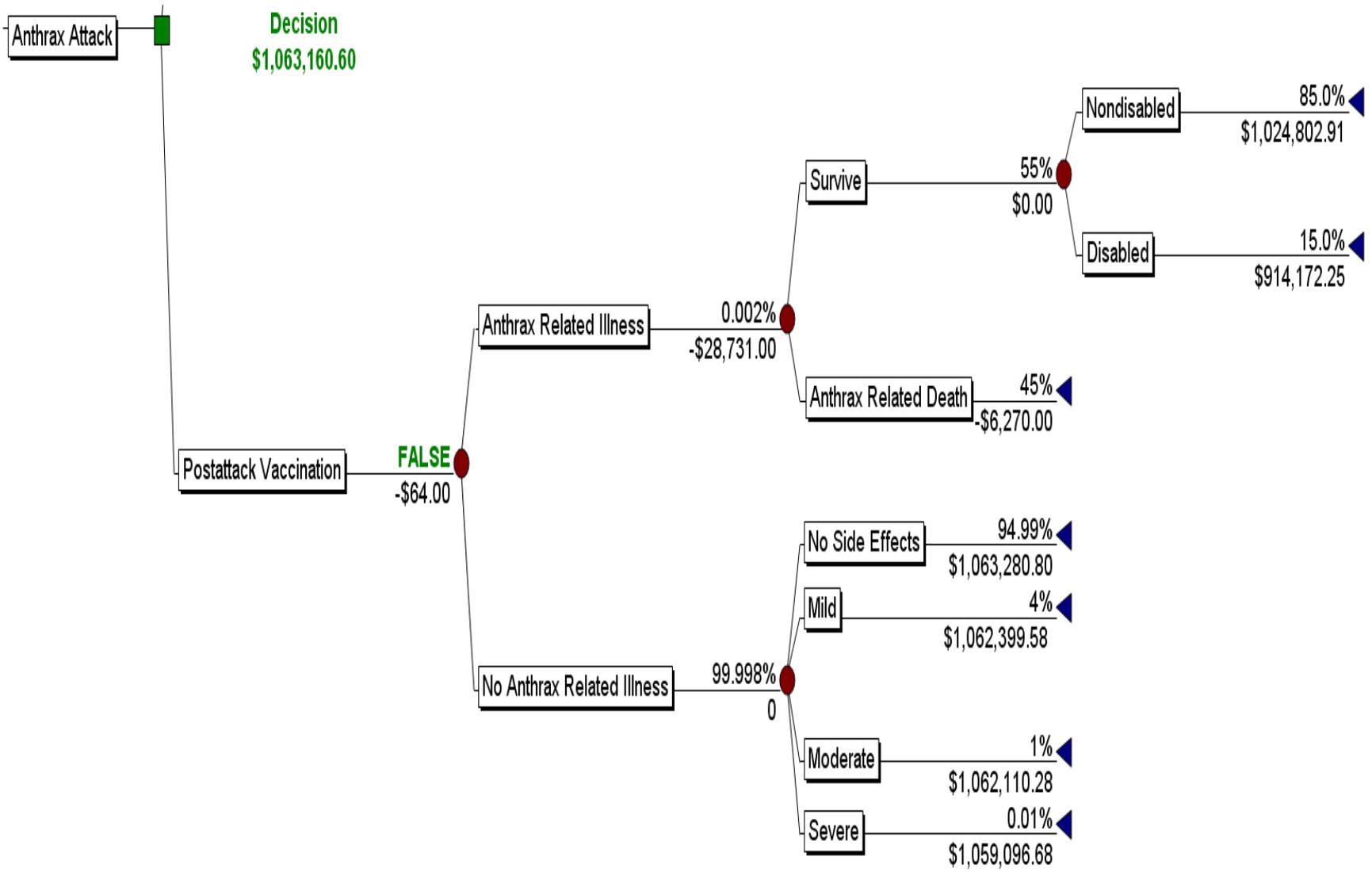
# Decision level for prospective anthrax risk

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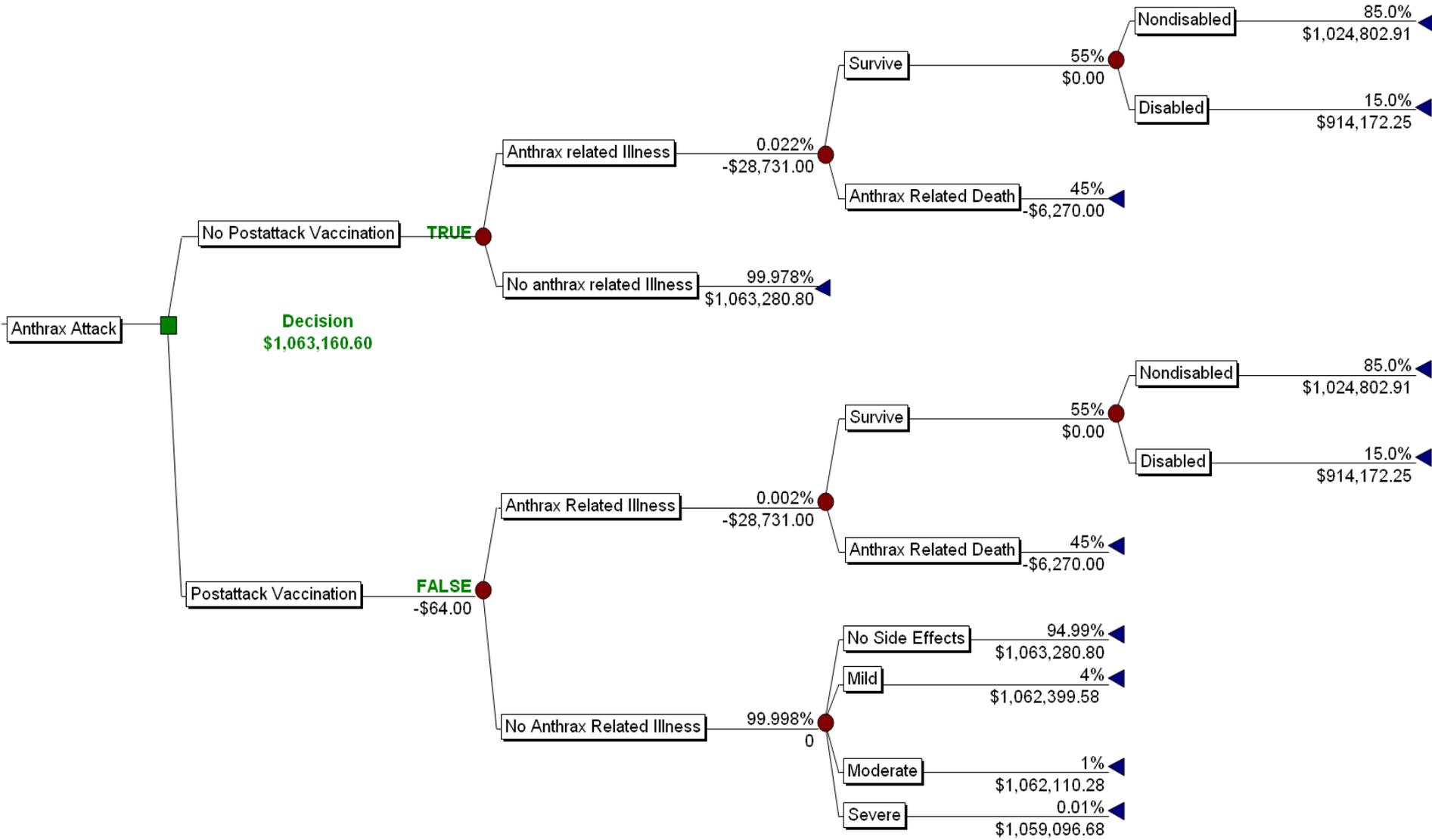
# Decision level for prospective anthrax risk

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# Decision level for prospective anthrax risk

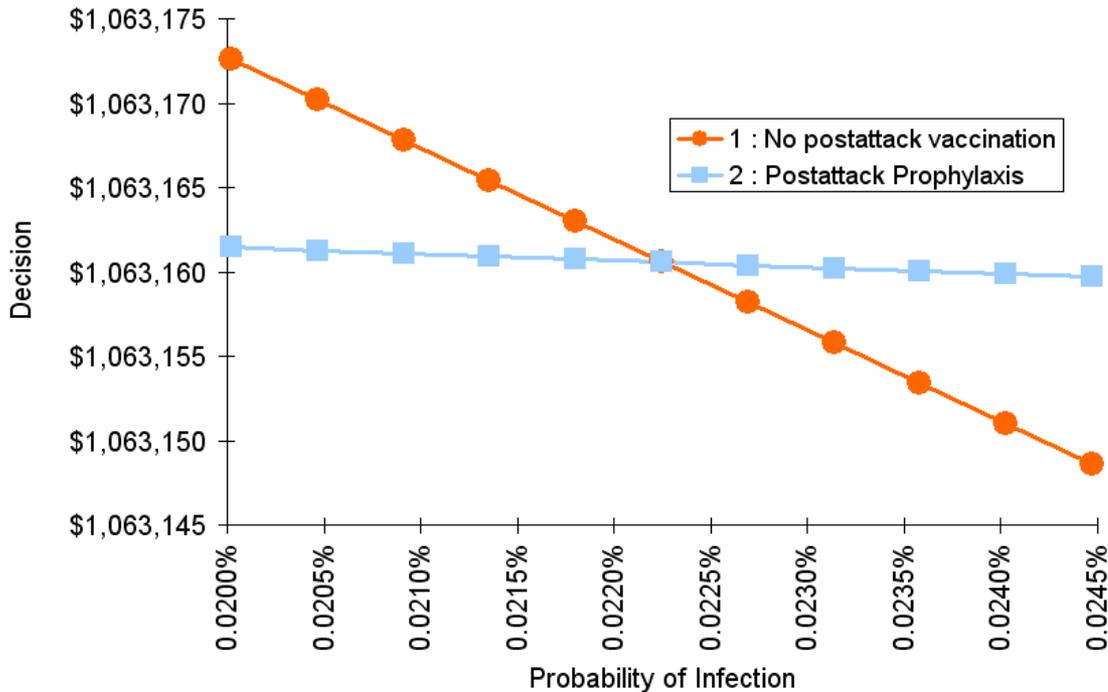
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# Decision level for prospective anthrax risk

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SWITCHOVER ANALYSIS

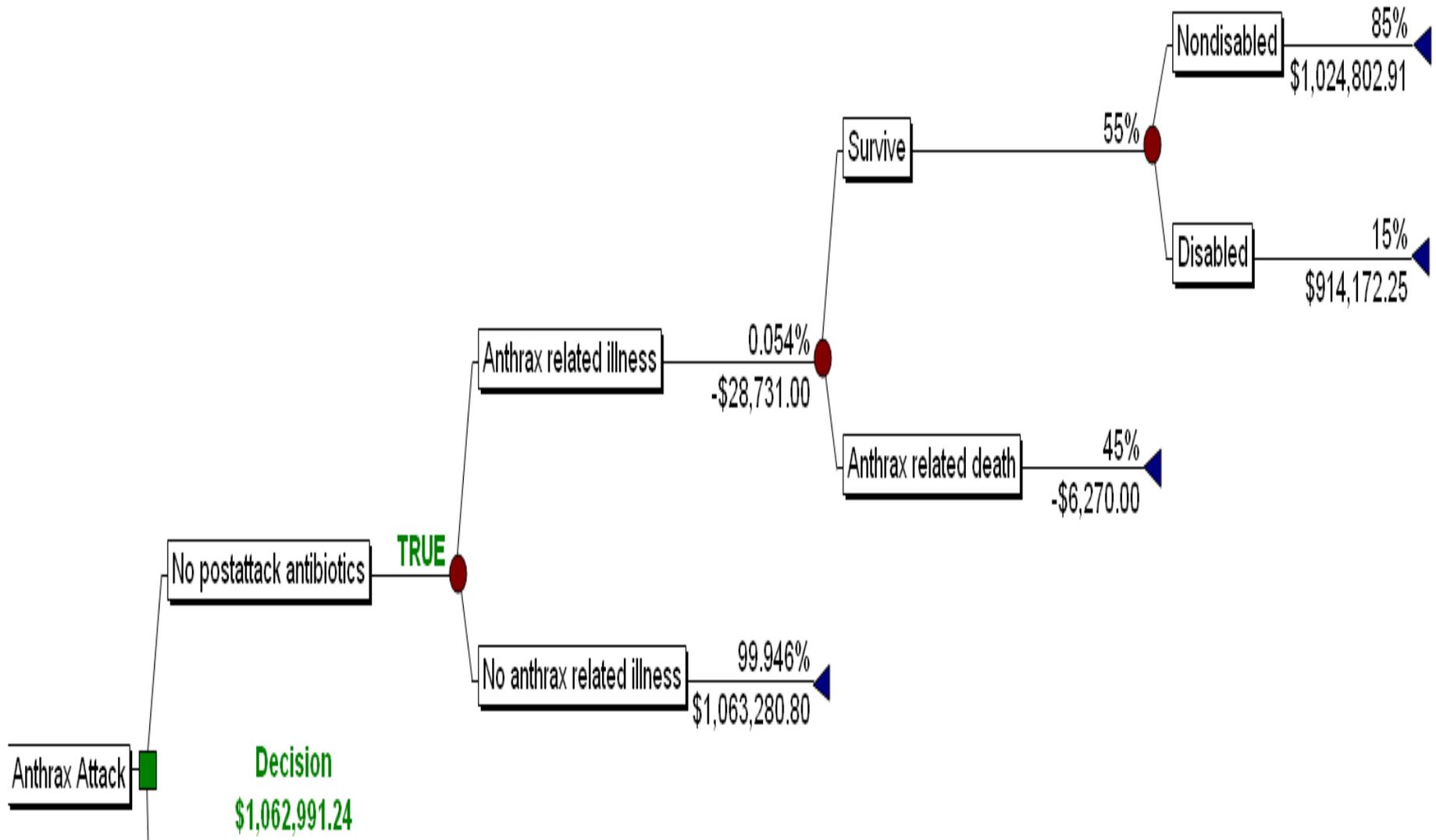


No action alternative is preferred for probability of infection < 0.022%, or 1 people in 4,495

This risk can be related to aerosol exposure using a dose-response function

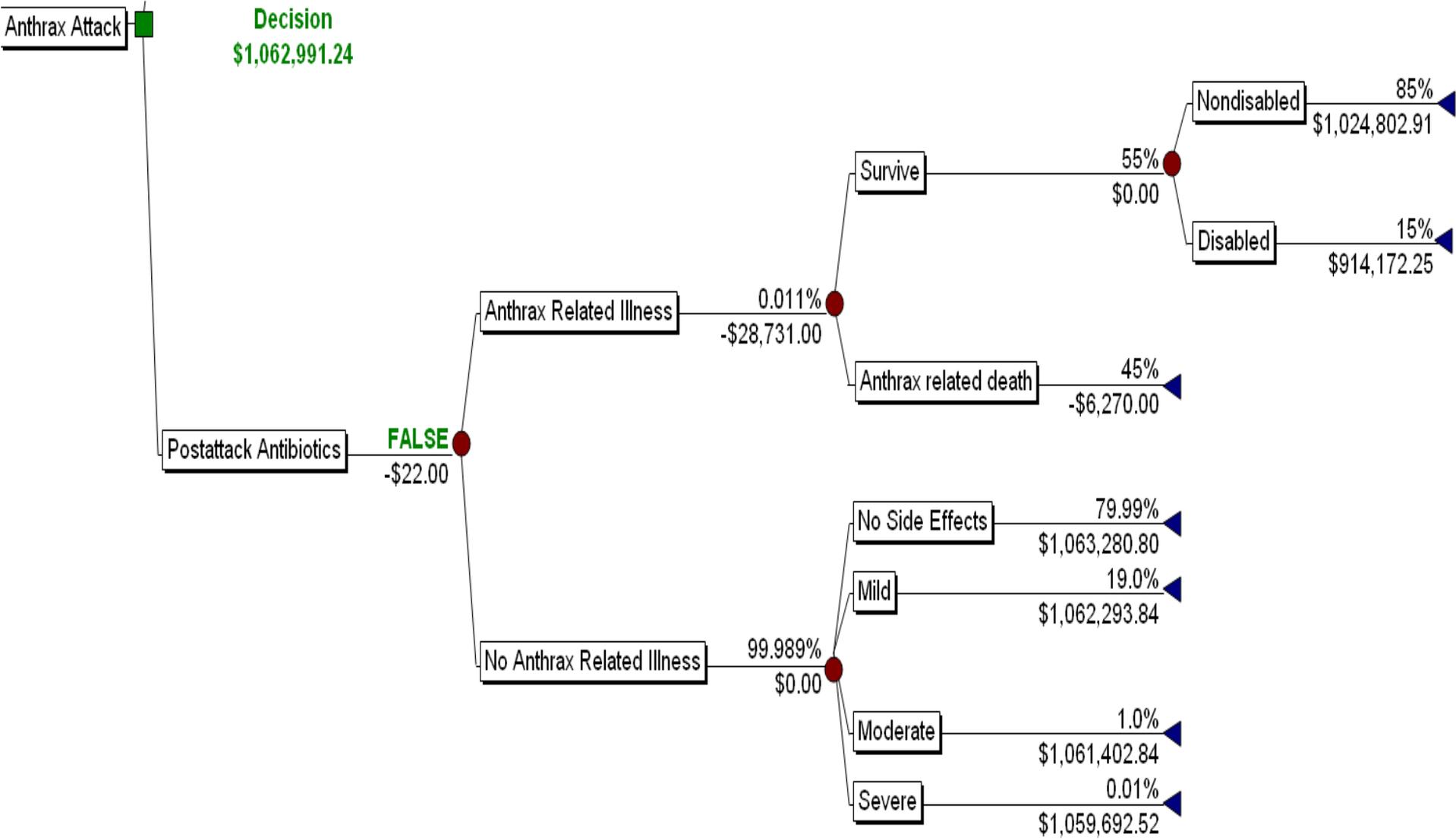
# Decision level for retrospective anthrax risk

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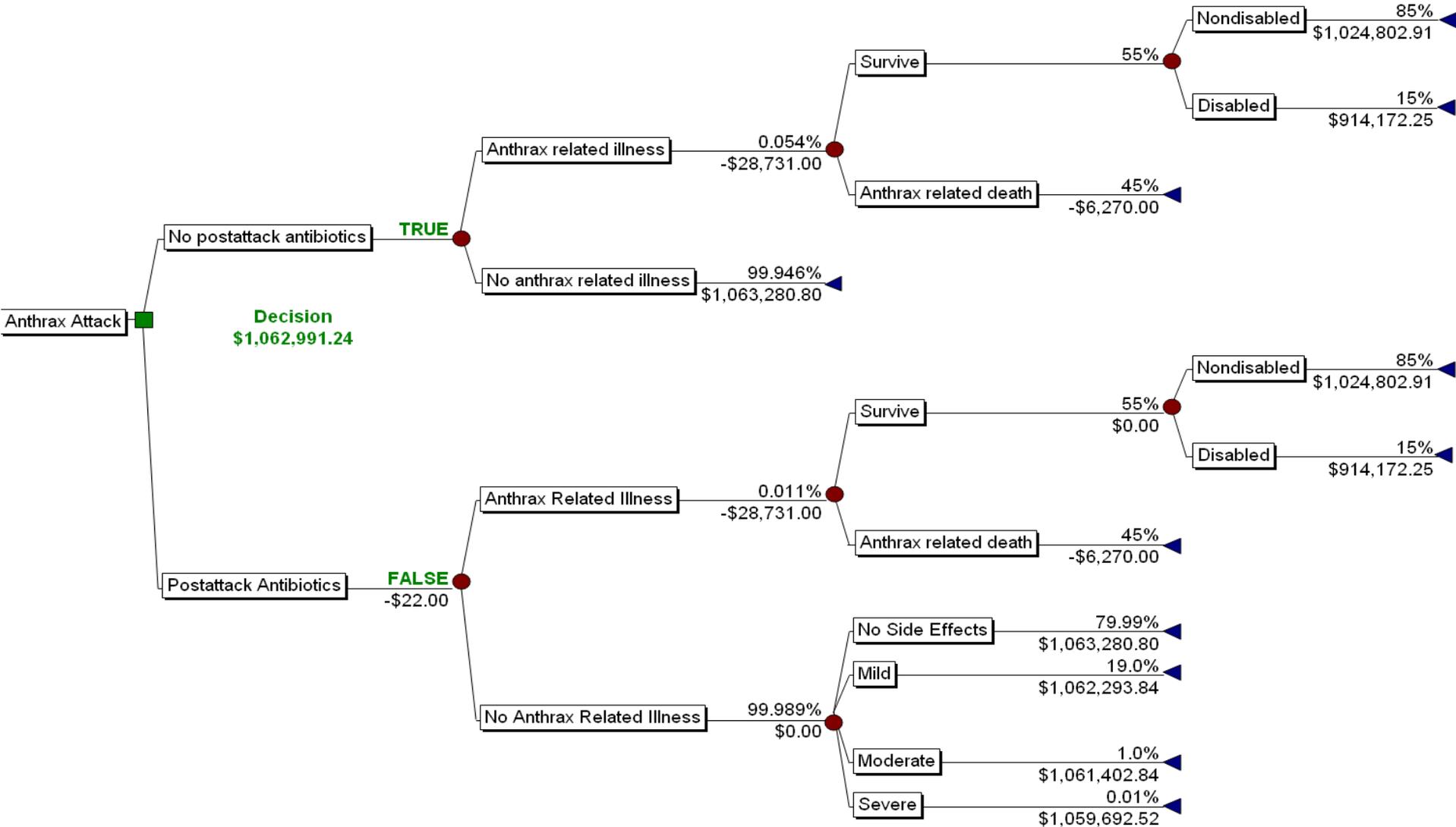
# Decision level for retrospective anthrax risk

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# Decision level for retrospective anthrax risk

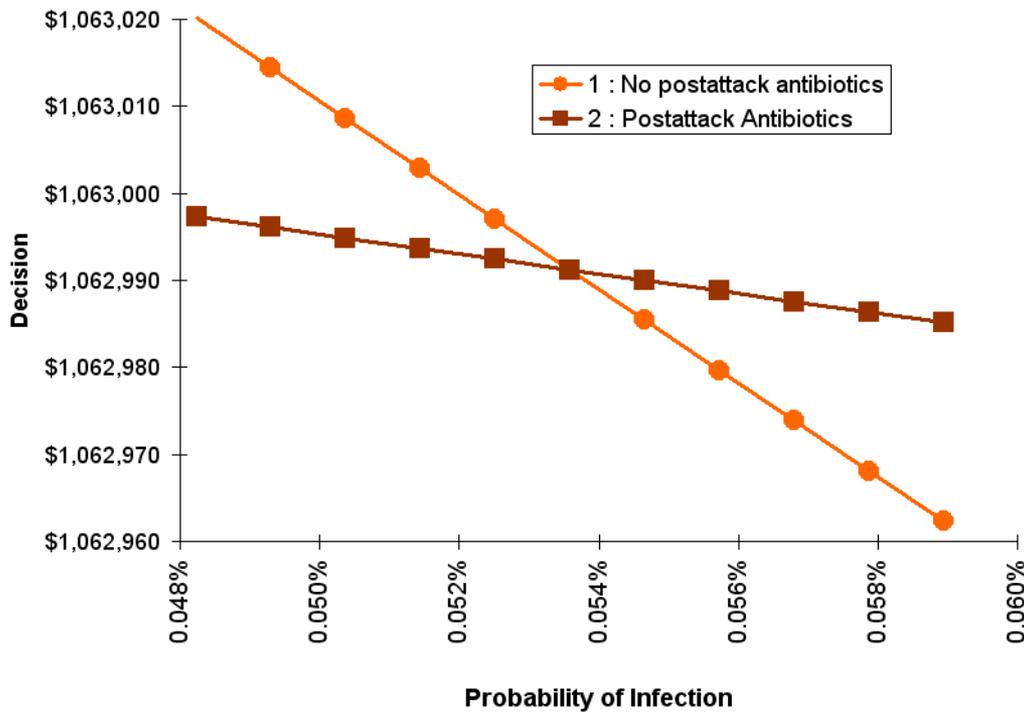
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# Decision level for retrospective anthrax risk

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SWITCHOVER ANALYSIS



No action alternative is preferred for probability of infection < 0.054%, or 1 people in 1,866

This risk can be related to aerosol exposure using a dose-response function

# Sensitivity Analysis

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	Prospective/ Vaccination	Retrospective/ Antibiotics
QALY=\$50K	1 in 4,495 0.022%	1 in 1,866 0.054%
QALY=\$100K	1 in 6,151 0.016%	1 in 1,917 0.052%
QALY=\$200K	1 in 7,620 0.013%	1 in 1,945 0.051%

# Conclusion

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- Benefit-cost analysis can suggest a level at which the decision to treat is justified
  - ▣ Many assumptions required about both empirical uncertainties and values
  - ▣ An expected-value analysis such as this may not be appropriate for all decision makers
- The conclusion will be sensitive to many factors including the value of a QALY

# Future Work

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- Sensitivity analysis for all uncertain model inputs
- Evaluation of the model for new treatments
- Consideration of the decision to remediate