

## Supplementary Online Content

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### **eReferences**

This supplementary material has been provided by the authors to give readers additional information about their work.



**eTable 1. Disease Progression Inputs Used in the CVD PREDICT Model**

| <b>Parameter</b>                                           | <b>Value</b>               | <b>Source</b>                                              |
|------------------------------------------------------------|----------------------------|------------------------------------------------------------|
| <b><i>From Disease Free State</i></b>                      |                            |                                                            |
| Non-CVD death                                              | Age and sex-specific table | NCHS 2010 <sup>1</sup>                                     |
| Stroke event                                               | Calibrated risk score      | Wolf 1991 <sup>2</sup>                                     |
| CHD event                                                  | Calibrated risk score      | Anderson 1991 <sup>3</sup>                                 |
| % CHD Cardiac Arrest                                       | Age and sex-specific table | Weinstein 1987 <sup>4</sup>                                |
| % CHD MI (male)                                            | 0.35                       | NHLBI 2006 <sup>5</sup>                                    |
| % CHD MI (female)                                          | 0.2                        | NHLBI 2006 <sup>5</sup>                                    |
| <b><i>From Stroke state</i></b>                            |                            |                                                            |
| Acute (1-year) risk of death                               | 0.15                       | Carandang 2006 <sup>6</sup>                                |
| Chronic (post 1 <sup>st</sup> -year) MI                    | 0.022                      | Touze 2005 <sup>7</sup>                                    |
| Chronic (post 1 <sup>st</sup> -year) stroke                | Calibrated risk score      | Wolf 1991 <sup>2</sup>                                     |
| <b><i>From MI state</i></b>                                |                            |                                                            |
| Acute (1-year) risk of death                               | 0.15                       | Mozaffarian 2016 <sup>8</sup>                              |
| Acute CABG                                                 | 0.082                      | Fang 2010 <sup>9</sup>                                     |
| Acute PTCA                                                 | 0.3                        | Fang 2010 <sup>9</sup>                                     |
| % Procedure death                                          | 0.0015                     | Williams 2006 <sup>10</sup>                                |
| Acute 2nd MI (no PTCA)                                     | 0.06                       | Capewell 2006 <sup>11</sup>                                |
| Acute 2nd MI (after PTCA)                                  | 0.053                      | Windecker 2014 <sup>12</sup>                               |
| Chronic (post 1 <sup>st</sup> -year) repeat MI             | 0.064                      | Jokhadar 2004 <sup>13</sup>                                |
| Chronic (post 1 <sup>st</sup> -year) repeat MI (with PTCA) | 0.052                      | Jokhadar 2004 <sup>13</sup> , Windecker 2014 <sup>12</sup> |

|                                                                |        |                                                             |
|----------------------------------------------------------------|--------|-------------------------------------------------------------|
| <b><i>From MI and CABG State</i></b>                           |        |                                                             |
| Acute post-CABG death                                          | 0.027  | Peterson 2004 <sup>14</sup>                                 |
| Acute 2nd MI                                                   | 0.047  | Windecker 2014 <sup>12</sup>                                |
| Repeat MI                                                      | 0.049  | Yusuf 1994 <sup>15</sup> , Windecker 2014 <sup>12</sup>     |
| <b><i>From Angina State</i></b>                                |        |                                                             |
| Acute (1-year) risk of death                                   | 0.045  | Capewell 2006 <sup>11</sup>                                 |
| Acute (1-year) risk of cardiac arrest                          | 0.006  | Hsia 2008 <sup>16</sup>                                     |
| Acute (1-year) risk of MI                                      | 0.035  | Hemingway 2003 <sup>17</sup>                                |
| Acute (1-year) risk of MI (with PTCA)                          | 0.031  | Hemingway 2003 <sup>17</sup> , Windecker 2014 <sup>12</sup> |
| Acute CABG                                                     | 0.2    | Ford 2007 <sup>18</sup>                                     |
| Acute PTCA                                                     | 0.3    | Ford 2007 <sup>18</sup>                                     |
| % Procedure Death                                              | 0.0015 | Assumption: same as MI                                      |
| Chronic (post 1 <sup>st</sup> -year) MI                        | 0.035  | Assumption: same as acute MI                                |
| Chronic (post 1 <sup>st</sup> -year) MI (with PTCA)            | 0.029  | Windecker 2014 <sup>12</sup>                                |
| <b><i>From Angina and CABG state</i></b>                       |        |                                                             |
| Acute post-CABG death                                          | 0.027  | Assumption: same as MI-CABG                                 |
| Acute 2nd MI                                                   | 0.028  | Windecker 2014 <sup>12</sup>                                |
| Chronic (post 1 <sup>st</sup> -year) MI                        | 0.0278 | Hemingway 2003 <sup>17</sup> ; Windecker 2014 <sup>12</sup> |
| <b><i>From Cardiac Arrest state</i></b>                        |        |                                                             |
| Acute (within 1 year) death                                    | 0.954  | Nichol 2008 <sup>19</sup>                                   |
| MI event                                                       | 0.064  | Assumption: same as MI                                      |
| <b><i>Chronic (post 1<sup>st</sup>-year) CVD Mortality</i></b> |        |                                                             |
| Proportion of chronic CVD deaths due to CVD                    | 0.28   | NHANES-based calculation                                    |
| Post-stroke all-cause mortality multiplier                     | 2.3    | Rosen 2010 <sup>20</sup>                                    |
| Post-CHD all-cause mortality multiplier (male)                 | 1.6    | Smolina 2012 <sup>21</sup>                                  |
| Post-CHD all-cause mortality multiplier                        | 2.1    | Smolina 2012 <sup>21</sup>                                  |

|                                                |     |                            |
|------------------------------------------------|-----|----------------------------|
| (female)                                       |     |                            |
| Post-CHD all-cause mortality >1 event (male)   | 3.4 | Smolina 2012 <sup>21</sup> |
| Post-CHD all-cause mortality >1 event (female) | 2.5 | Smolina 2012 <sup>21</sup> |

Note: see Pandya et al. 2017 for the full explanation of how these inputs were derived:

Pandya A, Sy S, Cho S, Alam S, Weinstein MC, Gaziano TA. Validation of a Cardiovascular Disease Policy Micro-simulation Model Using Both Survival and Receiver Operating Characteristic Curves. *Med Decis Making*. 2017 Oct;37(7):802-814.

**eTable 2. Utilities Used in the CVD PREDICT Model**

| <b>Parameter</b>                                                                        | <b>Base-case Value</b> | <b>Base-Case Source</b>                                 |
|-----------------------------------------------------------------------------------------|------------------------|---------------------------------------------------------|
| Disease Free                                                                            | 0.877                  | Sullivan 2006 (15), Mozafarran 2016 (3)                 |
| Chronic Cardiac Arrest                                                                  | 0.808                  | Sullivan 2006 <sup>22</sup> , Taylor 2009 <sup>23</sup> |
| Chronic MI                                                                              | 0.778                  | Sullivan 2006 <sup>22</sup>                             |
| Chronic MI with CABG                                                                    | 0.778                  | Sullivan 2006 <sup>22</sup>                             |
| Chronic Angina                                                                          | 0.768                  | Sullivan 2006 <sup>22</sup>                             |
| Chronic Angina with CABG                                                                | 0.768                  | Sullivan 2006 <sup>22</sup>                             |
| Chronic Stroke                                                                          | 0.768                  | Sullivan 2006 <sup>22</sup>                             |
| <i>Utilities for Acute Disease States (disabilities for acute state in parentheses)</i> |                        |                                                         |
| Acute Cardiac Arrest                                                                    | 0.770 (-0.0409)        | Sullivan 2006 <sup>22</sup>                             |
| Acute MI                                                                                | 0.737 (-0.0409)        | Sullivan 2006 <sup>22</sup>                             |
| Acute MI with CABG                                                                      | 0.737 (-0.0409)        | Sullivan 2006 <sup>22</sup>                             |
| Acute Angina                                                                            | 0.727 (-0.0412)        | Sullivan 2006 <sup>22</sup>                             |
| Acute Angina with CABG                                                                  | 0.727 (-0.0412)        | Sullivan 2006 <sup>22</sup>                             |
| Acute Stroke                                                                            | 0.716 (-0.0524)        | Sullivan 2006 <sup>22</sup>                             |
| <i>Disutilities for Events</i>                                                          |                        |                                                         |
| Repeat MI                                                                               | -0.041                 | Sullivan 2006 <sup>22</sup>                             |
| Repeat Stroke                                                                           | -0.052                 | Sullivan 2006 <sup>22</sup>                             |
| CABG                                                                                    | 0                      | assumption                                              |
| PTCA                                                                                    | 0                      | assumption                                              |
| Statin                                                                                  | -0.002                 | Gage 1996 <sup>24</sup> , Hutchins 2015 <sup>25</sup>   |
| Minor statin adverse event                                                              | -0.005                 | Lee 2010 <sup>26</sup> (2 days of life lost)            |

|                            |        |                                               |
|----------------------------|--------|-----------------------------------------------|
| Major statin adverse event | -0.038 | Lee 2010 <sup>26</sup> (2 weeks of life lost) |
|----------------------------|--------|-----------------------------------------------|

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**eTable 3. Costs (2017 US Dollars) Used in the CVD PREDICT Model**

| <b>Parameter</b>                                     | <b>Base-Case Value</b> | <b>Base-Case Source</b>                              |
|------------------------------------------------------|------------------------|------------------------------------------------------|
| <b><i>Costs for Chronic Disease States</i></b>       |                        |                                                      |
| Disease Free                                         | \$0                    | Assumption: None                                     |
| Chronic CHD                                          | \$3,368                | Lee 2010 <sup>26</sup>                               |
| Chronic Stroke                                       | \$2,225                | Pignone 2006 <sup>27</sup>                           |
| <b><i>Costs for Acute Disease States</i></b>         |                        |                                                      |
| Acute Cardiac Arrest                                 | \$20,277               | O'Sullivan 2011 <sup>28</sup>                        |
| Acute MI                                             | \$59,301               | O'Sullivan 2011 <sup>28</sup>                        |
| Acute Angina                                         | \$30,660               | O'Sullivan 2011 <sup>28</sup>                        |
| Acute Stroke                                         | \$20,127               | O'Sullivan 2011 <sup>28</sup>                        |
| <b><i>Costs for Procedures and Repeat Events</i></b> |                        |                                                      |
| Repeat MI                                            | \$59,301               | O'Sullivan 2011 <sup>28</sup>                        |
| Repeat Stroke                                        | \$20,127               | O'Sullivan 2011 <sup>28</sup>                        |
| CABG                                                 | \$38,797               | O'Sullivan 2011 <sup>28</sup>                        |
| PTCA                                                 | \$36,556               | O'Sullivan 2011 <sup>28</sup>                        |
| <b><i>Screening Costs</i></b>                        |                        |                                                      |
| Non-lab test (GP visit in Stage 1)                   | \$79                   | Pletcher 2009 <sup>29</sup>                          |
| Cholesterol (lab) test                               | \$37                   | Pletcher 2009 <sup>29</sup>                          |
| # extra GP visits during Stage 2                     | 1                      | Assumption                                           |
| # lab tests/year after treatment                     | 1                      | Lazar 2011 <sup>30</sup> , Expert Opinion            |
| # GP visits/year after treatment                     | 1                      | Lazar 2011 <sup>30</sup> , Expert Opinion            |
| <b><i>Statin Drug and Adverse Event Costs</i></b>    |                        |                                                      |
| Statin                                               | \$281                  | Redbook 2009 <sup>31</sup>                           |
| Anti-hypertensive                                    | \$217                  | Nuckols 2011 <sup>32</sup>                           |
| Aspirin                                              | \$8                    | Pignone 2006 <sup>27</sup>                           |
| ACE Inhibitor                                        | \$55                   | Shah 2011 <sup>33</sup> , Redbook 2009 <sup>31</sup> |



|                     |         |                                                      |
|---------------------|---------|------------------------------------------------------|
| Beta Blocker        | \$55    | Shah 2011 <sup>33</sup> , Redbook 2009 <sup>31</sup> |
| Mild adverse event  | \$188   | Lee 2010 <sup>26</sup>                               |
| Major adverse event | \$7,400 | Lee 2010 <sup>26</sup>                               |

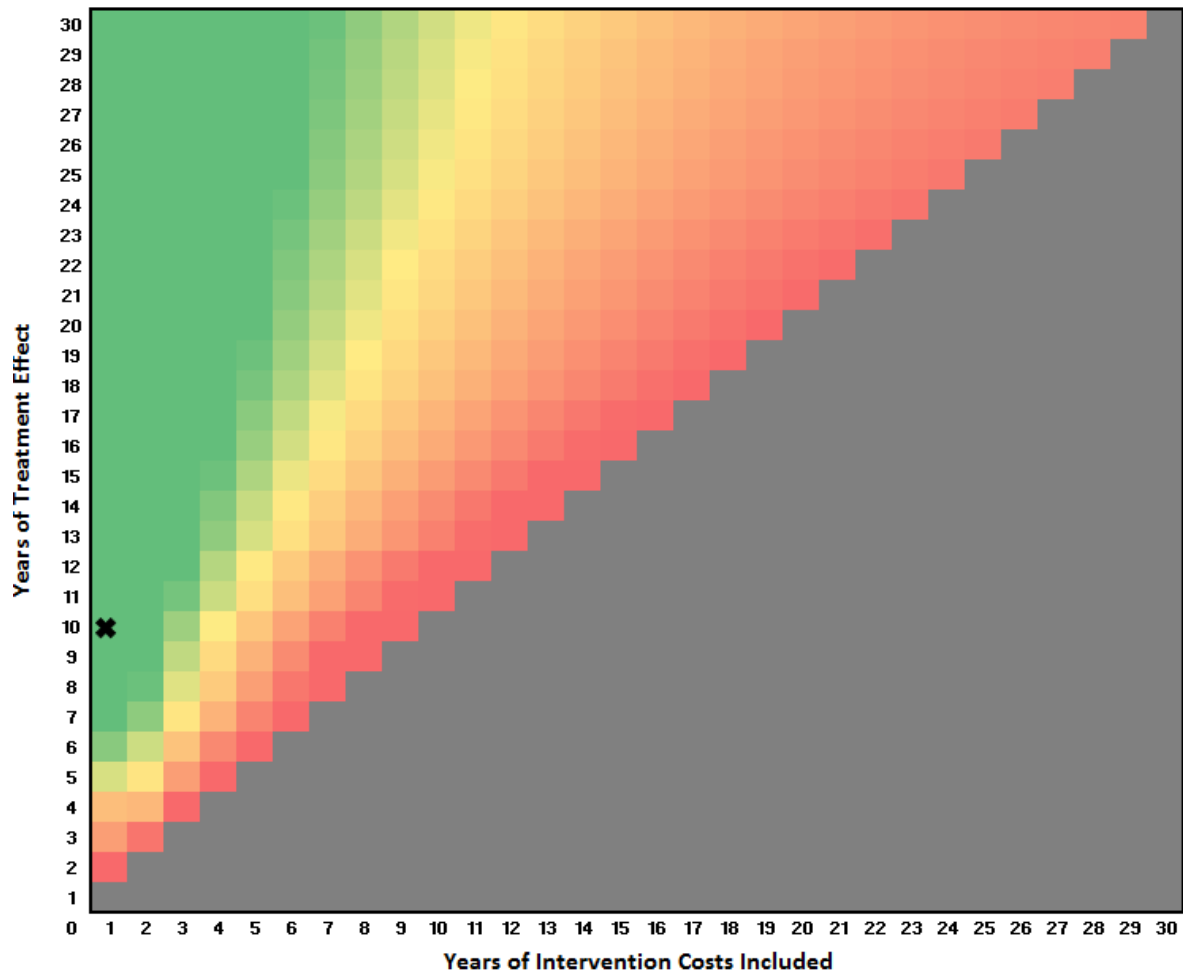
Note: see Pandya et al. 2017 for the full explanation of how these inputs were derived:

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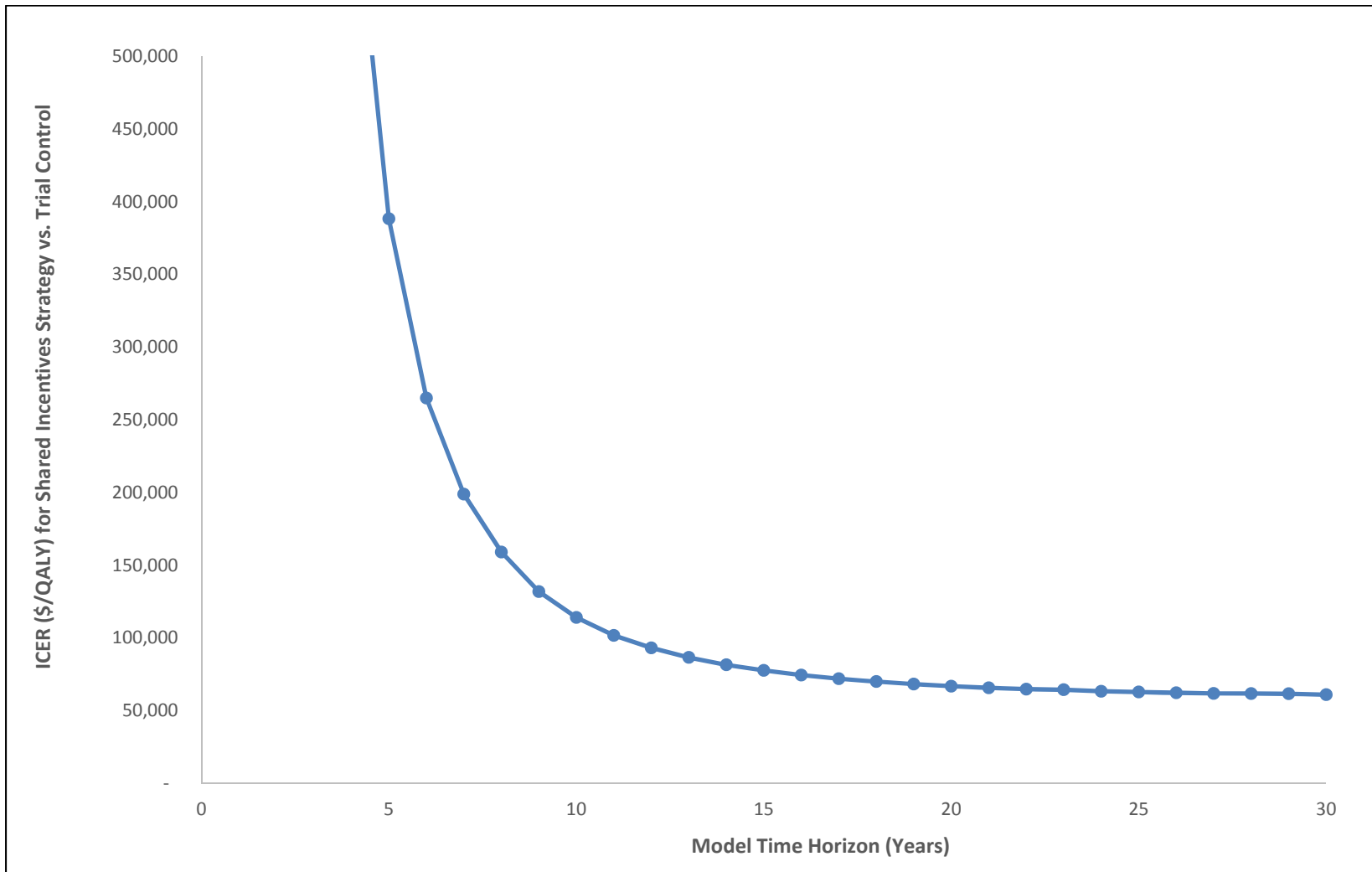
**eTable 4. Trial and Model Baseline Characteristics for Each Strategy (Treatment Arm)**

| <b>Characteristic</b>                   | <b>Trial</b> | <b>Model</b> |
|-----------------------------------------|--------------|--------------|
| n                                       | 1,503        | 1,000,000    |
| Age, mean (SD), years                   | 61.99 (8.7)  | 61.5 (11.9)  |
| Female sex (%)                          | 42.7         | 30.7         |
| African American (%)                    | 15.5         | 10.6         |
| Currently smoking (%)                   | *            | 31.8         |
| History of diabetes                     | 18.1         | 35.2         |
| Systolic blood pressure mean (SD), mmHg | 129.1 (14.9) | 136.6 (20.8) |
| Total cholesterol mean (SD), mg/dL      | *            | 228.5 (46.4) |
| LDL cholesterol mean (SD), mg/dL        | 160.6 (27.2) | 153.9 (39.3) |
| HDL cholesterol mean (SD), mg/dL        | *            | 44.3 (12.9)  |
| History of coronary heart disease (%)   | 34.5         | 7.7          |
| Taking cholesterol medication (%)       | 33.2         | 27.3         |
| CHD risk mean (SD)                      | 19.8 (8.7)   | 19.2 (8.4)   |

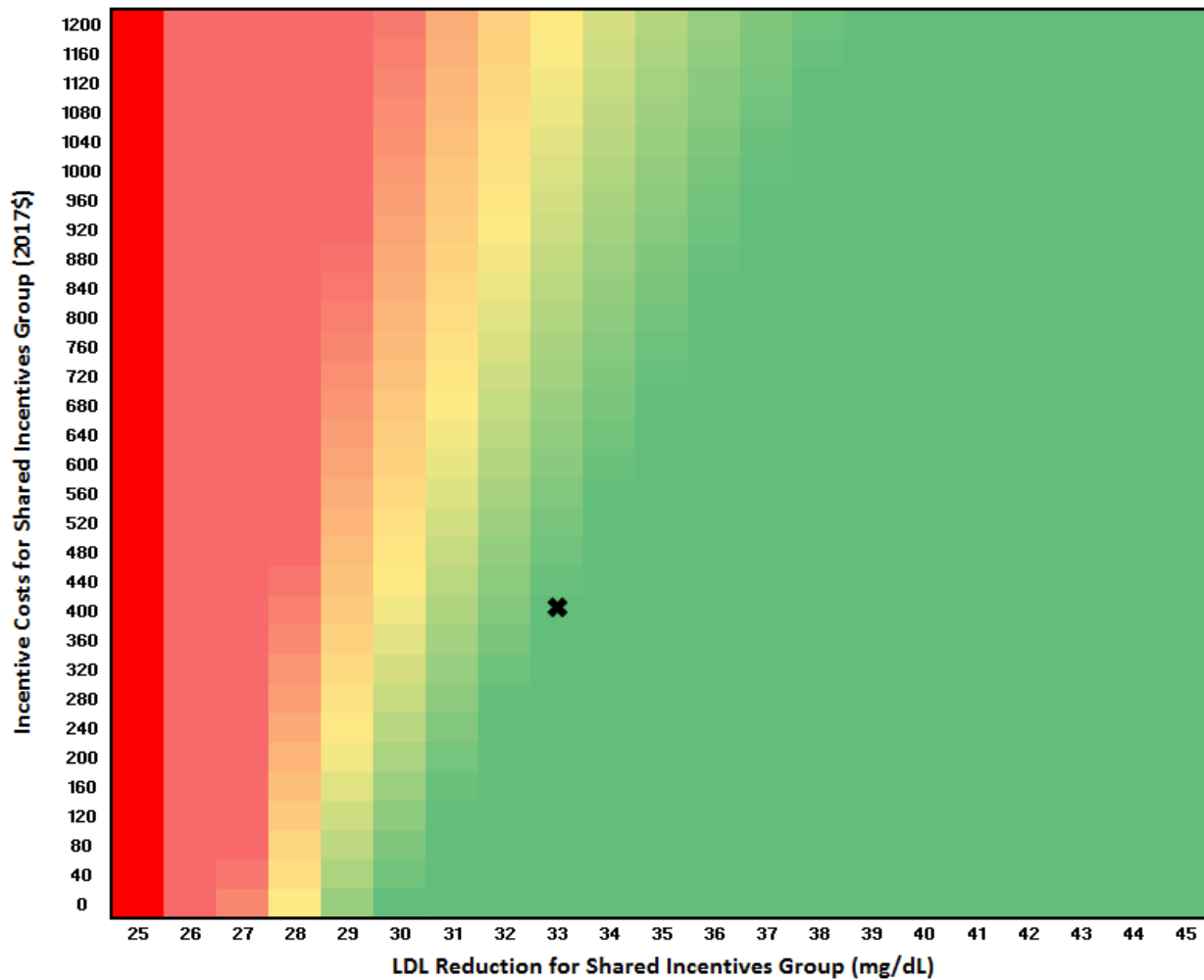
\*not included in trial dataset



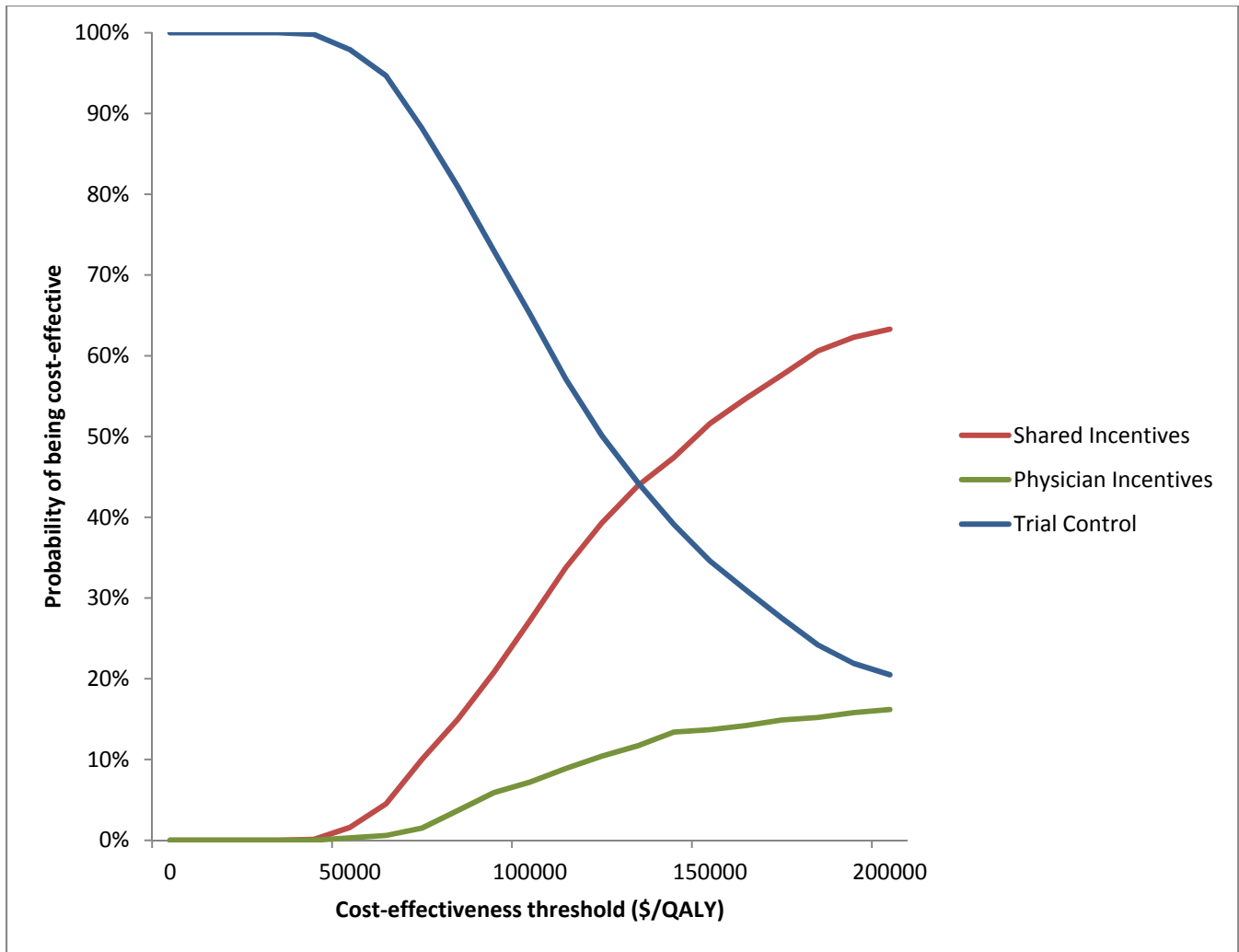
**eFigure 1. Two-Way Sensitivity Analysis Showing the ICER for the Shared Incentives Strategy Compared to the Trial Control for Different Combinations of LDL Reduction Waning and Years of Intervention Costs, Assuming a Cost-effectiveness Threshold of \$150,000/QALY.** The green regions show combinations of values that resulted in an ICER <\$100,000/QALY for the shared incentives strategy compared to the trial control strategy; yellow indicates an ICER of \$150,000/QALY, red indicates an ICER of >\$200,000/QALY; orange indicates an ICER between \$150,000-200,000/QALY; gray indicates implausible results (years where intervention costs are included but treatment effects are not observed in those years). “X” marks the base-case assumption and result (treatment effect linearly wanes to zero by year 10).



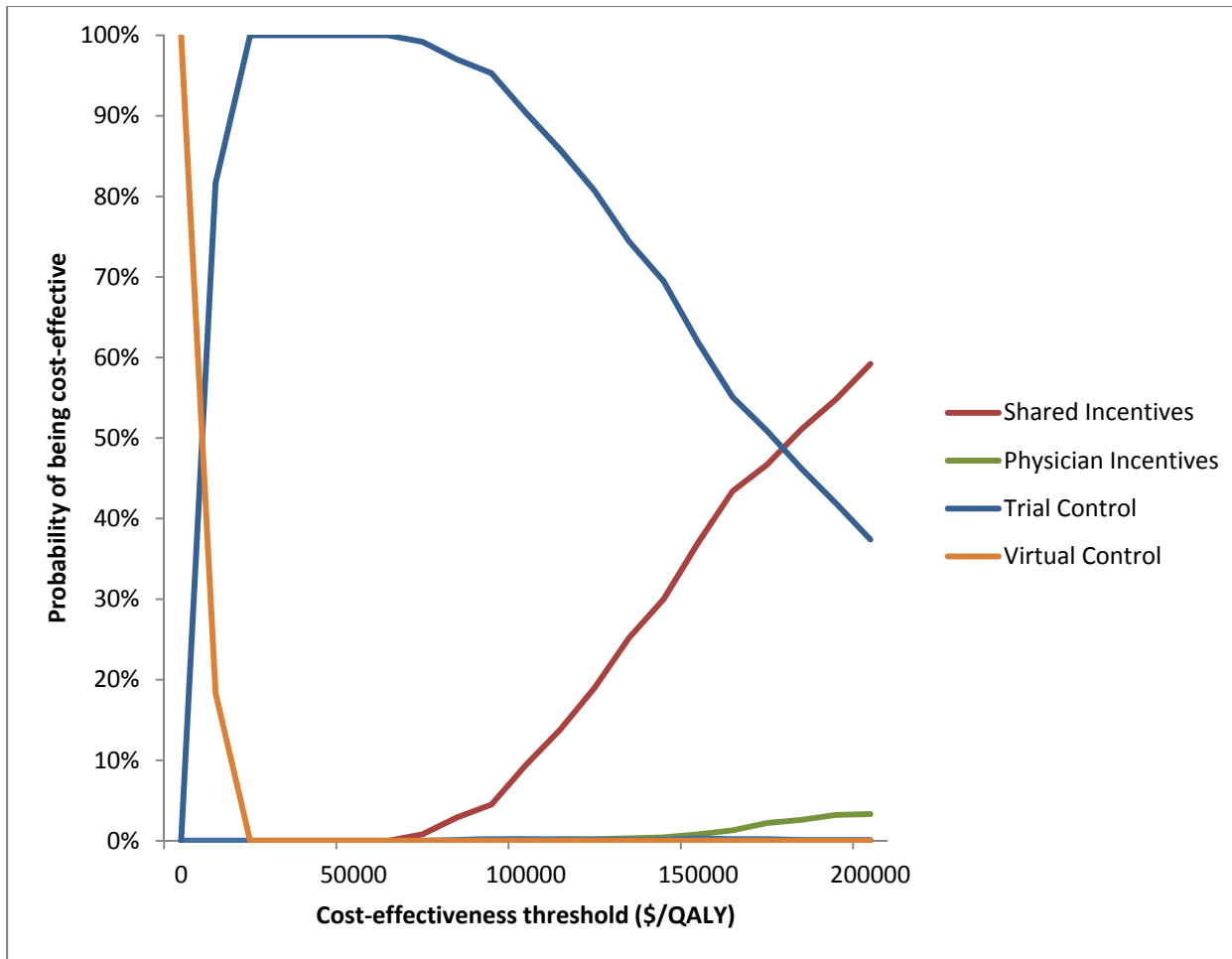
**eFigure 2. One-Way Sensitivity Analysis Showing the ICER for the Shared Incentives Strategy as a Function of Analytical Time Horizon.**



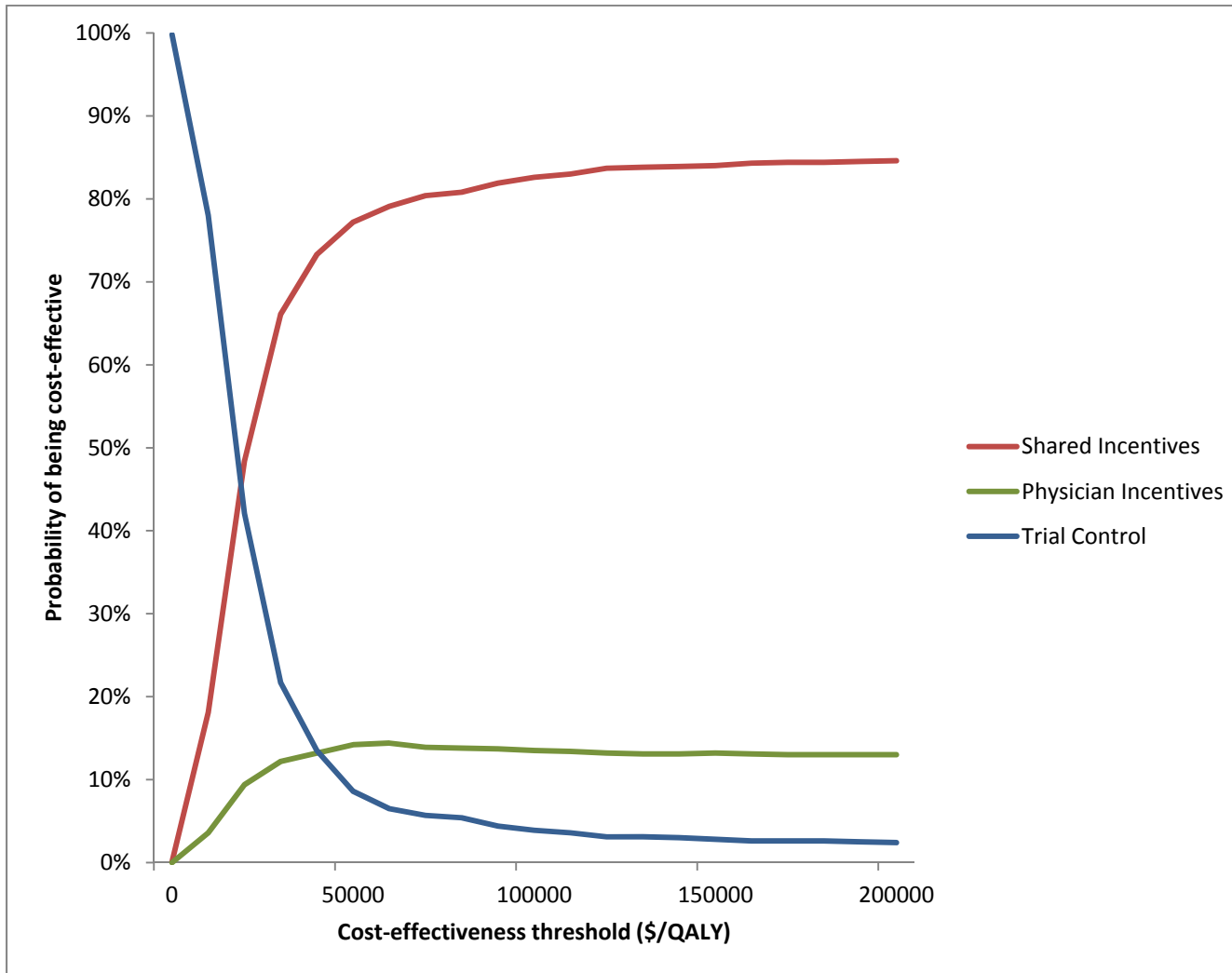
**eFigure 3. Two-Way Sensitivity Analysis Showing the ICER for the Shared Incentives Strategy Compared to the Trial Control for Different Combinations of LDL Cholesterol Reductions and Average Shared Financial Incentives Payouts.** The green regions show combinations of values that resulted in an ICER <\$50,000/QALY for the shared incentives strategy compared to the trial control strategy; yellow indicates an ICER of \$100,000/QALY, red indicates an ICER of >\$200,000/QALY; orange indicates an ICER between \$100,000-200,000/QALY; “X” marks the base-case assumption and result.



**eFigure 4. Cost-effectiveness Acceptability Curve (CEAC) for the Probabilistic Sensitivity Analysis (PSA) for Scenario of 5-Year LDL Reduction Waning.**

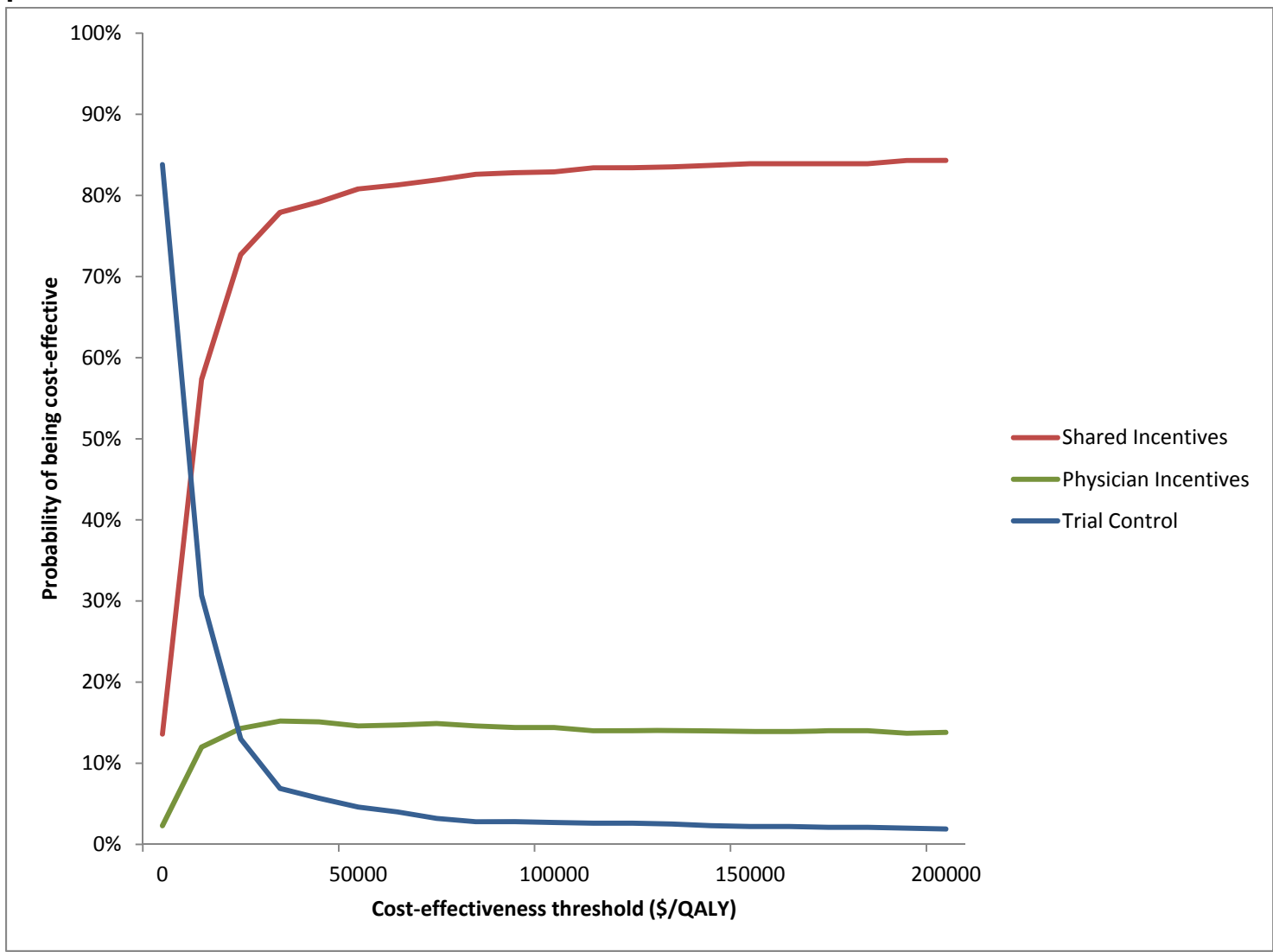


**eFigure 5. Cost-effectiveness Acceptability Curve (CEAC) for the Probabilistic Sensitivity Analysis (PSA) for Scenario of 10-Year LDL Reduction Waning and Including 5 Years of Intervention Costs.**



**eFigure 6. Cost-effectiveness Acceptability Curve (CEAC) for the Probabilistic Sensitivity Analysis (PSA) for Scenario of 30-Year LDL Reduction Waning**





**eFigure 7. Cost-effectiveness Acceptability Curve (CEAC) for the Probabilistic Sensitivity Analysis (PSA) for Scenario of Lifetime LDL Benefit Duration.**

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