

## COST-EFFECTIVENESS ANALYSIS

Health Economics Introductory Workshop  
Leonard Davis Institute of Health Economics

Sponsored by Janssen Pharmaceuticals, Inc.

Henry Glick, Ph.D.  
[www.uphs.upenn.edu/dgimhsr](http://www.uphs.upenn.edu/dgimhsr)

August 22, 2014



---

---

---

---

---

---

---

---

### Outline

- Introduction to cost-effectiveness analysis (CEA)
- Choice criteria for CEA



---

---

---

---

---

---

---

---

### Cost-Effectiveness Analysis (I)

- Estimates costs and outcomes of intervention
- Costs and outcomes expressed in different units
  - If outcomes aggregated using measures of preference (e.g., quality-adjusted life years saved), referred to as cost utility analysis
- Most used form of economic analysis



---

---

---

---

---

---

---

---

### Cost-Effectiveness Analysis (II)

- Results meaningful:
  - If there exists a predefined standard (i.e., a threshold, or maximum acceptable cost-effectiveness ratio, or an acceptability criterion) against which they can be compared
    - e.g., \$50,000 per year of life saved might be considered the threshold, or
  - Compared with other accepted and rejected interventions (e.g., against league tables)



---

---

---

---

---

---

---

---

### Cost-Effectiveness "History"

- \$/Life saved
- \$/Year of life saved (YOL)
- \$/Quality adjusted life year saved (QALY)
  
- ??? US Congress and outlawing QALYs ???



---

---

---

---

---

---

---

---

### Cost-Effectiveness Ratios

- Cost-effectiveness ratio
$$\frac{\text{Costs}_1 - \text{Costs}_2}{\text{Effects}_1 - \text{Effects}_2}$$
- A ratio exists for every pair of options
  - 1 option (case series), no ratios calculated
  - 2 options, 1 ratio
  - 3 options, 3 ratios (option 1 versus option 2, option 1 versus option 3, and option 2 versus option 3)
- In "efficient" selection algorithm, don't necessarily calculate all possible ratios



---

---

---

---

---

---

---

---

### Colorectal Cancer Screening

- Suppose 5 screening strategies have the following discounted costs and life expectancies:

Treatment	Cost	YOLS
S1 Sig Q10	1290	17.378
S2 Sig Q5	1535	17.387
S3 U+Sig, Q10	1810	17.402
S4 C, Q10	2030	17.396
S5 U+Sig, Q5	2035	17.407

Frazier AL, et al. JAMA. 2000;284:1954-61.

- What calculations might help us make a choice between them?




---

---

---

---

---

---

---

---

---

---

### Mistake #1

- Divide a therapy's cost by its outcome; compare resulting ratios

Treatment	Cost		YOLS	=	C/Y
S1 Sig Q10	1290	÷	17.378	=	74.23
S2 Sig Q5	1535	÷	17.387	=	88.28
S3 U+Sig, Q10	1810	÷	17.402	=	104.01
S4 C, Q10	2030	÷	17.396	=	116.69
S5 U+Sig, Q5	2035	÷	17.407	=	116.91

- Sometimes mistakenly referred to as the average cost-effectiveness ratios




---

---

---

---

---

---

---

---

---

---

### Dividing a Therapy's Costs by Its Effects is "Generally Uninformative"

	Cost	Effect	Ratio
Example 1			
Rx1	2,800	0.28	10,000
Rx2	5,800	0.29	20,000
Example 2			
Rx1	2,800	0.28	10,000
Rx2	11,200	0.56	20,000




---

---

---

---

---

---

---

---

---

---

Dividing a Therapy's Costs by Its Effects is  
"Generally Uninformative"

	Cost	Effect	Ratio
Example 1			
Rx1	2,800	0.28	10,000
Rx2	5,800	0.29	20,000
	$(5,800-2,800) / (0.29-0.28) = 300,000$		
Example 2			
Rx1	2,800	0.28	10,000
Rx2	11,200	0.56	20,000
	$(11,200-2,800) / (0.56-0.28) = 30,000$		




---

---

---

---

---

---

---

---

---

---

Mistake #2

- Calculate ratios for all therapies versus S1; compare resulting ratios

Treatment	Cost	ΔCost	YOLS	ΔYOLS	ACER
S1 Sig Q10	1290	--	17.378	--	--
S2 Sig Q5	1535	245	17.387	.009	27,222
S3 U+Sig, Q10	1810	520	17.402	.024	21,667
S4 C, Q10	2030	740	17.396	.018	41,111
S5 U+Sig, Q5	2035	745	17.407	.029	25,690

- Correctly referred to as average cost-effectiveness ratios




---

---

---

---

---

---

---

---

---

---

Average Cost-Effectiveness Ratio

- If these were the correct ratios, what should we conclude?

Treatment	Cost	ΔCost	YOLS	ΔYOLS	ACER
S1 Sig Q10	1290	--	17.378	--	--
S2 Sig Q5	1535	245	17.387	.009	27,222
S3 U+Sig, Q10	1810	520	17.402	.024	21,667
S4 C, Q10	2030	740	17.396	.018	41,111
S5 U+Sig, Q5	2035	745	17.407	.029	25,690




---

---

---

---

---

---

---

---

---

---

### What is Good Value?

- The “cost-effective” strategy delivers the largest health outcome that we are still willing to pay for
- Why don't the average ratios provide this information?




---

---

---

---

---

---

---

---

### What's Wrong with the Average Cost-Effectiveness Ratio

Treatment	Cost	ΔCost	YOLS	ΔYOLS	ACER
S1 Sig Q10	1290	--	17.378	--	--
S3 U+Sig, Q10	1810	520	17.402	.024	21,667
S5 U+Sig, Q5	2035	745	17.407	.029	25,690

- 25,690 for U+Sig, Q5 gives credit for the 520 we are already spending and the .024 YOLs we are already receiving from S3
- Compared to S3, we are spending almost 50% more for S5 and receiving only about 20% more of the outcome




---

---

---

---

---

---

---

---

### Incremental Cost-Effectiveness Ratio

- Basic idea for correct ratio: calculate ratio for S2 vs S1, S3 vs S2, S4 vs S3 and S5 vs S4

Treatment	Cost	ΔCost	YOLS	ΔYOLs	ICER
S1 Sig Q10	1290	--	17.378	--	--
S2 Sig Q5	1535	245	17.387	.009	27,222
S3 U+Sig, Q10	1810	275	17.402	.015	18,333
S4 C, Q10	2030	220	17.396	-.006	-36,667
S5 U+Sig, Q5	2035	5	17.407	.011	455

- But not quite right




---

---

---

---

---

---

---

---

Problem 1

- Never want to spend more and obtain less outcome as in S4 vs S3. **S4 is strongly dominated by S3**

Treatment	Cost	$\Delta$	YOLS	$\Delta$	ICER
S1 Sig Q10	1290	--	17.378	--	--
S2 Sig Q5	1535	245	17.387	.009	27,222
<b>S3 U+Sig, Q10</b>	<b>1810</b>	<b>275</b>	<b>17.402</b>	<b>.015</b>	<b>18,333</b>
<b>S4 C, Q10</b>	<b>2030</b>	<b>220</b>	<b>17.396</b>	<b>-.006</b>	<b>-36,667</b>
S5 U+Sig, Q5	2035	225	17.407	.005	45,000

- S4 should be eliminated from consideration for adoption




---

---

---

---

---

---

---

---

---

---

---

---

Problem 2

- Don't want to buy less outcome for a higher cost per unit of outcome as in S2 vs S3: **S2 weakly dominated by S3**

Treatment	Cost	$\Delta$	YOLS	$\Delta$	ICER
S1 Sig Q10	1290	--	17.378	--	--
<b>S2 Sig Q5</b>	<b>1535</b>	<b>245</b>	<b>17.387</b>	<b>.009</b>	<b>27,222</b>
<b>S3 U+Sig, Q10</b>	<b>1810</b>	<b>275</b>	<b>17.402</b>	<b>.015</b>	<b>18,333</b>
S4 C, Q10	2030	220	17.396	-.006	S-Dom
S5 U+Sig, Q5	2035	225	17.407	.005	45,000




---

---

---

---

---

---

---

---

---

---

---

---

Problem 2

- S2 should be eliminated from consideration for adoption  
– Must recalculate ratio for S3 vs S1

Treatment	Cost	$\Delta$	YOLS	$\Delta$	ICER
S1 Sig Q10	1290	--	17.378	--	--
S2 Sig Q5	1535	245	17.387	.009	W-Dom
S3 U+Sig, Q10	1810	520	17.402	.024	21,667
S4 C, Q10	2030	220	17.396	-.006	S-Dom
S5 U+Sig, Q5	2035	225	17.407	.005	45,000




---

---

---

---

---

---

---

---

---


---

---

---

**Steps for Calculating ICER**

- Step 1: Rank order therapies in ascending order of either costs or outcomes (final ordering of nondominated therapies unaffected by variable chosen)
  - Already correctly ordered by cost
- Step 2: Eliminate therapies that are strongly dominated (i.e., have increased costs and reduced effects compared with at least one other alternative)
  - S4 is strongly dominated by S3
- Step 3: Compute incremental cost-effectiveness ratios for each adjacent pair of remaining outcomes (e.g., between options 1 and 2; between options 2 and 3; etc.)




---

---

---

---

---


---

---

---

**Steps for Calculating ICER (2)**

- If resulting ratios are ranked from lowest to highest, can skip to step 6. If not...
- Step 4: Eliminate therapies that are less effective (costly) but have a higher cost-effectiveness ratio than next higher ranked therapy (weakly dominated/extended dominance)
  - “S2 is weakly dominated by S3”; “eliminate S2 because of extended dominance by S3”




---

---

---

---

---


---

---

---

**Steps for Calculating ICER (2)**

- Step 5: Recalculate ratio for next higher ranked therapy vs next lower ranked therapy
  - E.g., S3 vs S1
  - Recalculated ratio will always be higher than original ratio, but can't be higher than weakly dominated ratio
    - E.g.,  $27,222 > 21,666 > 18,333$
  - If resulting ratios still not ranked from lowest to highest, may need to repeat evaluation of weakly dominated therapies several times
    - After S2 is eliminated, ratios are ordered from lowest to highest
- Step 6: Identify acceptable ratio




---

---

---

---

---

---

---

---

### Reduced Table

- Candidates for adoption include S1, S3, and S5

Treatment	Cost	$\Delta$	YOLS	$\Delta$	ICER
S1 Sig Q10	1290	--	17.378	--	--
S3 U+Sig, Q10	1810	520	17.402	.024	21,667
S5 U+Sig, Q5	2035	225	17.407	.005	45,000

- If  $W < 21667$ , adopt S1
- If  $W \geq 21,667$  and  $< 45,000$ , adopt S3
- If  $W \geq 45,000$ , adopt S5




---

---

---

---

---

---

---

---

### Simultaneous Comparison

- Description of selection algorithm may suggest a path through different options, with adoption of lower cost/effect pairs before adoption of higher cost/effect pairs
- Not true
  - Selection algorithm is simply step-by-step procedure that simultaneously compares all options




---

---

---

---

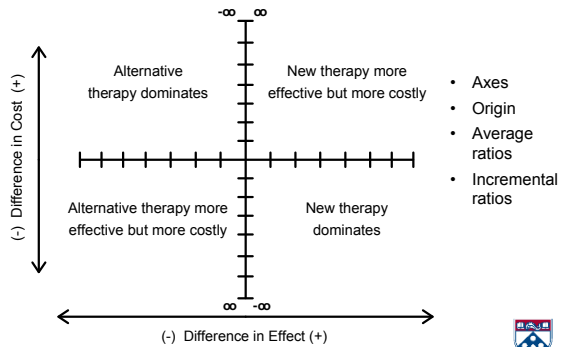
---

---

---

---

### Cost-Effectiveness Plane




---

---

---

---

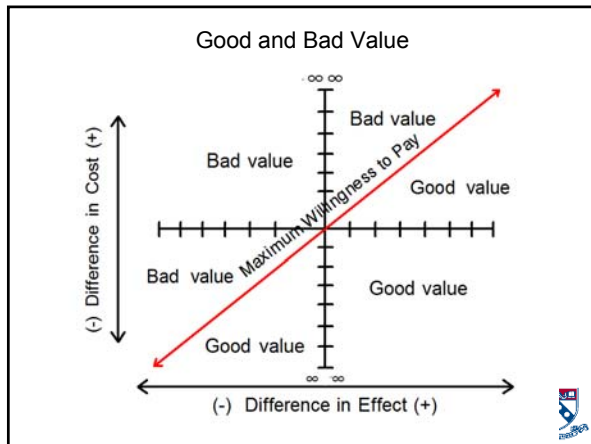
---

---

---

---






---

---

---

---

---

---

---

---

- ### What Is Maximum Acceptable Ratio?
- US Gov't
    - EPA: 9.1 M / life (~222K / undiscounted YOLS)
    - FDA: 7.9 M / life (~176K / undiscounted YOLS)
    - DOT: 6 M / life (~133K / undiscounted YOLS)
  - Australia: \$AU 42K - 76K /YOLS
  - Italy: €60,000/QALY
  - Netherlands: €80 000/QALY
  - Sweden: SEK 500,000 (€54,000) / QALY
  - UK: £20 - 30K / QALY
  - WHO report: 3 times GDP per DALY




---

---

---

---

---

---

---

---

- ### Take Home Messages
- Decision making using cost-effectiveness ratios requires attention to incremental cost-effectiveness ratios
  - To make decisions using these ratios, they must be compared to:
    - A predefined standard (i.e., an acceptability criterion) against which they can be compared (e.g., \$50,000 per year of life saved might be considered largest acceptable ratio), or
    - Other accepted and rejected interventions (e.g., against league tables)




---

---

---

---

---

---

---

---