

# Cost-Effectiveness Analysis of Infliximab versus Cyclosporine in Steroid-Refractory Acute Severe Ulcerative Colitis

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## Background

- Acute severe ulcerative colitis (ASUC) affects 25% of all ulcerative colitis (UC) patients, requiring hospitalization and treatment with intravenous steroids<sup>1</sup>
- In ASUC cases that are steroid-refractory, infliximab (INFLX) or cyclosporine (CsA) are indicated to induce remission, with the goal of therapy being avoidance of colectomy<sup>2</sup>
- There are no recent cost-effectiveness analyses comparing these two agents from a United States payer perspective utilizing data from multiple clinical trials<sup>3</sup>

## Objective

- Estimate the cost-effectiveness of infliximab compared with cyclosporine for steroid-refractory ASUC in U.S. adults from a commercial payer perspective

## Methods

- A decision tree was constructed (Figure 1; Table 1).
- Probability, cost, and utility inputs were derived from publicly available literature and resources (Table 2). Cost and utility were discounted at a 3% annual rate.
  - The main outcome measure was incremental cost per quality-adjusted life year (QALY) gained
  - For each annual time step, patients could remain in remission or undergo colectomy, with possible complications and death associated with colectomy
- Uncertainty was assessed through a one-way deterministic sensitivity analysis and scenario analysis.
  - Scenario 1: One year timeframe
  - Scenario 2: Two year timeframe
  - Scenario 3: Use of an infliximab biosimilar (infliximab-dybb)

TABLE 1: Summary of Key Model Characteristics

Population	Adults hospitalized with steroid-refractory ASUC
Comparators	Infliximab vs. cyclosporine
Perspective	U.S. commercial payer
Time Horizon	3 years

## Methods (Cont.)

FIGURE 1: Decision Tree Model

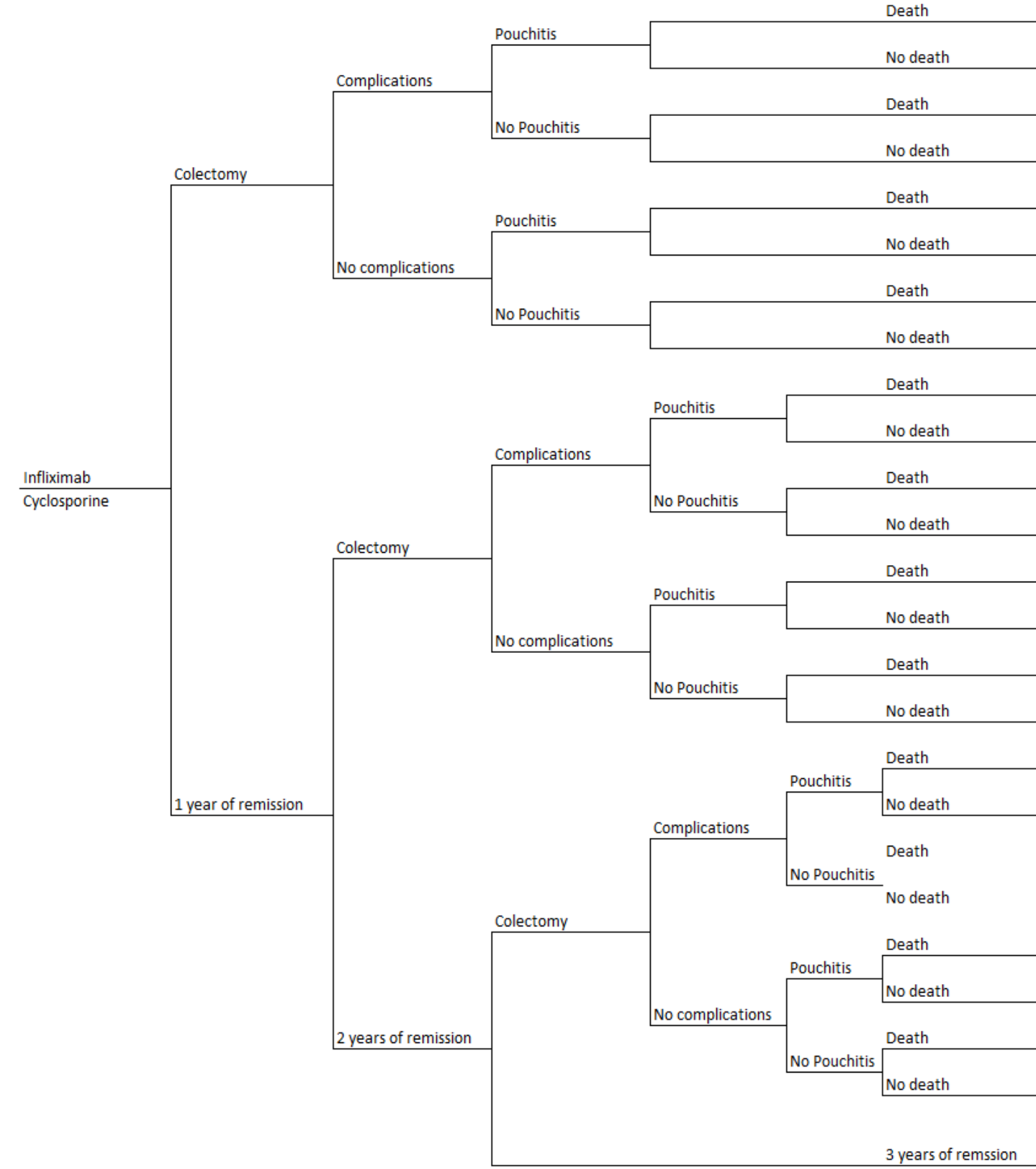


TABLE 2: Summary of Key Model Inputs

Probabilities	
Infliximab 3 year colectomy-free survival <sup>4</sup>	0.955
Cyclosporine 3 year colectomy-free survival <sup>4</sup>	0.939
UC colectomy complication (early/chronic pouchitis) <sup>5</sup>	0.213 / 0.155
UC colectomy death (emergent/elective) <sup>6</sup>	0.115 / 0.004
Mean Annual Costs (2023 US\$)	
Infliximab total drug cost (base/biosimilar) <sup>7, 8</sup>	\$4,194 / \$2,044
Cyclosporine total drug cost <sup>7,8,9</sup>	\$1,557
Colectomy <sup>10</sup>	\$41,400 <sup>10</sup>
Management of early complications <sup>5</sup>	\$1,2056
Management of chronic pouchitis <sup>11</sup>	\$21,873
Management in clinical remission and post colectomy <sup>12</sup>	\$5,153
Utilities	
1 year of remission <sup>13</sup>	0.81
1 year of post-colectomy <sup>14</sup>	0.79
1 year of post-colectomy with early complications <sup>15</sup>	0.49
1 year of post-colectomy with pouchitis <sup>15</sup>	0.40

## Results

TABLE 3: Base Case Results

	INFLX	CsA	Incremental Value
Cost	\$33,756	\$36,754	-\$2,989
Utility (QALY)	2.086	2.033	0.052
ICER (\$/QALY)	CsA is dominated		

TABLE 4: Scenario Analysis Results

Scenario 1		INFLX	CsA	Incremental Value
	Cost	\$17,721	\$22,230	-\$4,502
	Utility (QALY)	0.739	0.800	-0.060
Scenario 2	ICER (\$/QALY)	\$74,789		
	Cost (\$)	\$27,558	\$29,984	-\$2,424
	Utility (QALY)	1.434	1.614	-0.180
Scenario 3	ICER (\$/QALY)	\$13,438		
	Cost (\$)	\$31,614	\$36,754	-\$5,139
	Utility (QALY)	2.086	2.033	0.052
	ICER (\$/QALY)	CsA is dominated		

## Key Takeaways

INFLX dominated CsA in the base case and the third scenario, and was **found to be cost-effective in all scenarios** based on a \$150,000 USD willingness-to-pay.

Results were most sensitive to **utility** derived from disease remission and the post-colectomy state, as well as medication and colectomy costs.

Though INFLX was found to be less costly due to higher probability of colectomy-free survival, payoff from increased survival and accrual of QALYs was not observed until the **third year** post-treatment.

## Limitations

- Heterogeneity in study design and maintenance regimens
- Inputs were derived from studies in which differences in maintenance therapy were not accounted for and heterogenous

## References

1. Du L, Ha C. Epidemiology and Pathogenesis of Ulcerative Colitis. *Gastroenterol Clin North Am*. 2020;49(4):643-654. doi:10.1016/j.gtc.2020.07.005

2. Burri E, Maillard MH, Schoepfer AM, et al. Treatment Algorithm for Mild and Moderate-to-Severe Ulcerative Colitis: An Update. *Digestion*. 2020;101 Suppl 1:2-15. doi:10.1159/000504092

3. Alam MF, Longo M, Cohen D, et al. Infliximab versus cyclosporin in steroid resistant acute severe ulcerative colitis: a model-based cost-utility analysis of data from CONSTRUCT pragmatic trial. *BMC Health Serv Res*. 2023;23(1):226. Published 2023 Mar 8. doi:10.1186/s12913-023-09233-w

4. Szemes K, Sós A, Hegyi P, et al. Comparable Long-Term Outcomes of Cyclosporine and Infliximab in Patients With Steroid-Refractory Acute Severe Ulcerative Colitis: A Meta-Analysis. *Front Med (Lausanne)*. 2020;6:338. Published 2020 Jan 21. doi:10.3389/fmed.2019.00338

5. Zogg CK, Najjar P, Diaz AJ, et al. Rethinking Priorities: Cost of Complications After Elective Colectomy. *Ann Surg*. 2016;264(2):312-322. doi:10.1097/SLA.0000000000001511

6. Hajiravala L, Leonard C, Oraniglo G, Davis K, Barton J. Urgent Inpatient Colectomy Carries a Higher Morbidity and Mortality Than Elective Surgery. *J Surg Res*. 2021;268:394-404. doi:10.1016/j.jss.2021.06.081

7. CMS. Payment Allowance Limits for Medicare Part B Drugs Effective through October 31, 2023. 2023. <https://www.cms.gov/apps/ama/license.asp?file=https%3A/www.cms.gov/files/zip/October-2023-asg-pricing-life.zip>. Accessed November 13, 2023.

8. CMS. Physician Fee Schedule. 2023; <https://www.cms.gov/apps/physician-feeschedule/overview.aspx>. Accessed November 13, 2023.

9. IBM Micromedex RedBook. 2023. Accessed November 13, 2023.

10. Xu F, Liu Y, Wheaton AG, Rabarison KM, Croft JB. Trends and Factors Associated with Hospitalization Costs for Inflammatory Bowel Disease in the United States. *Appl Health Econ Health Policy*. 2019;17(1):77-91. doi:10.1007/s40258-018-0432-4

11. Barnes EL, Kappelman MD, Zhang X, Long MD, Sandler RS, Herfarth HH. Patients With Pouchitis Demonstrate a Significant Cost Burden in the First Two Years After Ileal Pouch-Anal Anastomosis. *Clin Gastroenterol Hepatol*. 2022;20(12):2908-2910.e2.

12. Cohen R, Skup M, Ozbay AB, et al. Direct and indirect healthcare resource utilization and costs associated with ulcerative colitis in a privately-insured employed population in the US. *J Med Econ*. 2015;18(6):447-456. doi:10.3111/j386698.2015.1021353

13. Gibson PR, Vazzey C, Black CM, et al. Relationship between disease severity and quality of life and assessment of health care utilization and cost for ulcerative colitis in Australia: a cross-sectional, observational study. *J Crohns Colitis*. 2014;8(7):598-606. doi:10.1016/j.crohns.2013.11.017

14. Brown C, Gibson PR, Hart A, et al. Long-term outcomes of colectomy surgery among patients with ulcerative colitis. *Springerplus*. 2015;4:573. Published 2015 Oct 5. doi:10.1186/s40064-015-1350-7

15. Arseneau KO, Sultan S, Provenzale DT, et al. Do patient preferences influence decisions on treatment for patients with steroid-refractory ulcerative colitis? *Clin Gastroenterol Hepatol*. 2006;4(9):1135-1142. doi:10.1016/j.cgh.2006.05.003

