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Immunotherapy for dMMR Endometrial Cancer: A Cost-Effectiveness Analysis

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Objectives:

Few treatment options exist for patients with advanced or recurrent endometrial cancer (EC) that has progressed after treatment with platinum-based therapy.¹ Dostarlimab and pembrolizumab are approved for patients with advanced cancers that have not responded to prior therapy and are mismatch repair deficient (dMMR).^{2,3} **This analysis aimed to add to the evidence on the cost-effectiveness of immunotherapy for advanced or recurrent dMMR EC.**

Methods:

- TREATMENTS: Immunotherapy (pembrolizumab or dostarlimab) versus usual care (pegylated doxorubicin, PLD)
- TIME HORIZON: 25 years
- PERSPECTIVE: Healthcare sector
- METHOD: Partitioned survival model to estimate probabilities of progression free survival, overall survival and death. State probabilities derived from survival curves in the published literature.^{4,5}
- COSTS AND UTILITIES: Derived from published literature and government estimates (Table 1). Expressed in 2022 US\$
- DISCOUNT RATE: 3%

References:
1. Rütten H, Verhoef C, van Weelden WJ, et al. Recurrent endometrial cancer: Local and systemic treatment options. *Cancers* (Basel). 2021;13(24):6275. doi:10.3390/cancers13246275
2. FDA grants accelerated approval to dostarlimab-gxly for dMMR advanced solid tumors. August 2021. US Food & Drug Administration. Accessed December 5, 2022. doi:10.3390/cancers13246275
3. FDA approves pembrolizumab for advanced endometrial carcinoma. March 2022. US Food & Drug Administration. Accessed December 5, 2022.
4. O'Malley DM, Bariani GM, Cassier PA, et al. Pembrolizumab in patients with microsatellite instability-high advanced endometrial cancer: Results from the KEYNOTE-158 study. *J Clin Oncol*. 2022;40(7):752-761. doi:10.1200/JCO.21.01874
5. Mathews C, Lorusso D, Coleman RL, Boklage S, Garside J. An indirect comparison of the efficacy and safety of dostarlimab and doxorubicin for the treatment of advanced and recurrent endometrial cancer. *Oncologist*. 2022;27(12):1058-1066. doi:10.1093/oncolo/oyac188
6. O'Malley DM, Bariani GM, Cassier PA, et al. Health-related quality of life with pembrolizumab monotherapy in patients with previously treated advanced microsatellite instability-high/mismatch repair deficient endometrial cancer in the KEYNOTE-158 study. *Gynecol Oncol*. 2022;166(2):245-253. doi:10.1016/j.ygyno.2022.06.005
7. October 2022 Quarterly Average Sales Price (ASP) Medicare Part B Drug Pricing Files and Revisions to Prior Quarterly Pricing Files. 2022. Accessed November 18, 2022.
8. Oaknin A, Gilbert L, Tinker AV, et al. Safety and antitumor activity of dostarlimab in patients with advanced or recurrent DNA mismatch repair deficient/microsatellite instability-high (dMMR/MSI-H) or proficient/stable (MMRp/MSS) endometrial cancer: interim results from GARNET-a phase I, single-arm study. *J Immunother Cancer*. 2022;10(1). doi:10.1136/jitc-2021-003777
9. Kalidindi Y, Jung J, Feldman R. Differences in spending on provider-administered chemotherapy by site of care in Medicare. *Am J Manag Care*. 2018;24(7):328-333.
10. CMS. Physician Fee Schedule—CY 2020 Physician Fee Schedule Final Rule. 2022. Available from: <https://www.cms.gov/Medicare/Medicare-Fee-for-Service-Payment/PhysicianFeeSched>
11. Bekelman JE, Halpern SD, Blankart CR, et al. Comparison of site of death, health care utilization, and hospital expenditures for patients dying with cancer in 7 developed countries. *JAMA*. 2016;315(3):272-283. doi:10.1001/jama.2015.18603
12. Kebede N, Shah R, Shah A, Corman S, Nwankwo C. Treatment patterns and economic burden among cervical and endometrial cancer patients newly initiating systemic therapy. *Future Oncol*. 2022;18(8):953-964. doi:10.2217/fo-2021-0772

Key Conclusions

- Dostarlimab and pembrolizumab led to increases in life expectancy and quality of life compared to PLD.
- Compared to pembrolizumab, dostarlimab resulted in more QALYs gained, at a lower cost per QALY.
- Based on current prices, these therapies would not be considered cost-effective at a willingness-to-pay threshold of \$150,000/QALY.
- More work is needed to understand clinical trajectories following treatment with immunotherapy and cycles needed to attain maximum benefit.

Table 1: Key Model Inputs

Transition Probabilities	
Drug	Distribution
Doxorubicin	
Progression-free survival	Log-normal (Mean=1.68, SD=0.98)
Overall survival	Log-normal (Mean=2.58, SD=0.94)
Pembrolizumab	
Progression-free survival	Log-normal (Mean=2.58, SD=1.97)
Overall survival	Log-logistic (Shape=1.19, Scale=40.92)
Dostarlimab	
Progression-free survival	Log-normal (Mean=2.71, SD=1.83)
Overall survival	Log-normal (Mean=3.78, SD=1.68)
Utilities	
Description	Base Value (Lower, Upper)
Stable disease	0.72 (0.62, 0.82) ⁶
Progressed disease	0.63 (0.43, 0.83) ⁶
Costs	
Description	Base Value (Lower, Upper)
Doxorubicin	\$1,680 (\$1,344, \$2,016) ^{5,7}
Pembrolizumab	\$10,910 (\$8,728, \$13,092) ^{4,7}
Dostarlimab, first 4 cycles	\$11,038 (\$8,830, \$13,245) ^{7,8}
Dostarlimab, subsequent cycles	\$22,075 (\$17,660, \$26,490) ^{7,8}
Medication administration	\$198 (\$82, \$284) ⁹
Physician visit	\$130 ¹⁰
Cost of care in final year of life	\$11,047 (\$10,909, \$11,184) ¹¹
Per-cycle cost of third line therapy	\$12,916 (\$0, \$46,820) ¹²

Results:

- Pembrolizumab was extended dominated by dostarlimab (Table 2).
- Neither immunotherapy was cost-effective at a willingness-to-pay threshold of \$150,000 per QALY (Table 2).
- Results were most sensitive to the costs of third line therapy following progression and maximum cycles of immunotherapy administered (Figure 1).

Table 2: Base Case Results

	Life Years	QALYs	Costs	ICER
Doxorubicin (PLD)	1.15	0.77	\$101,121	
Pembrolizumab	4.33	2.97	\$713,246	\$278,123
Dostarlimab	4.99	3.39	\$804,941	\$268,910

Figure 1A: One-Way Sensitivity Analysis: Pembrolizumab vs PLD

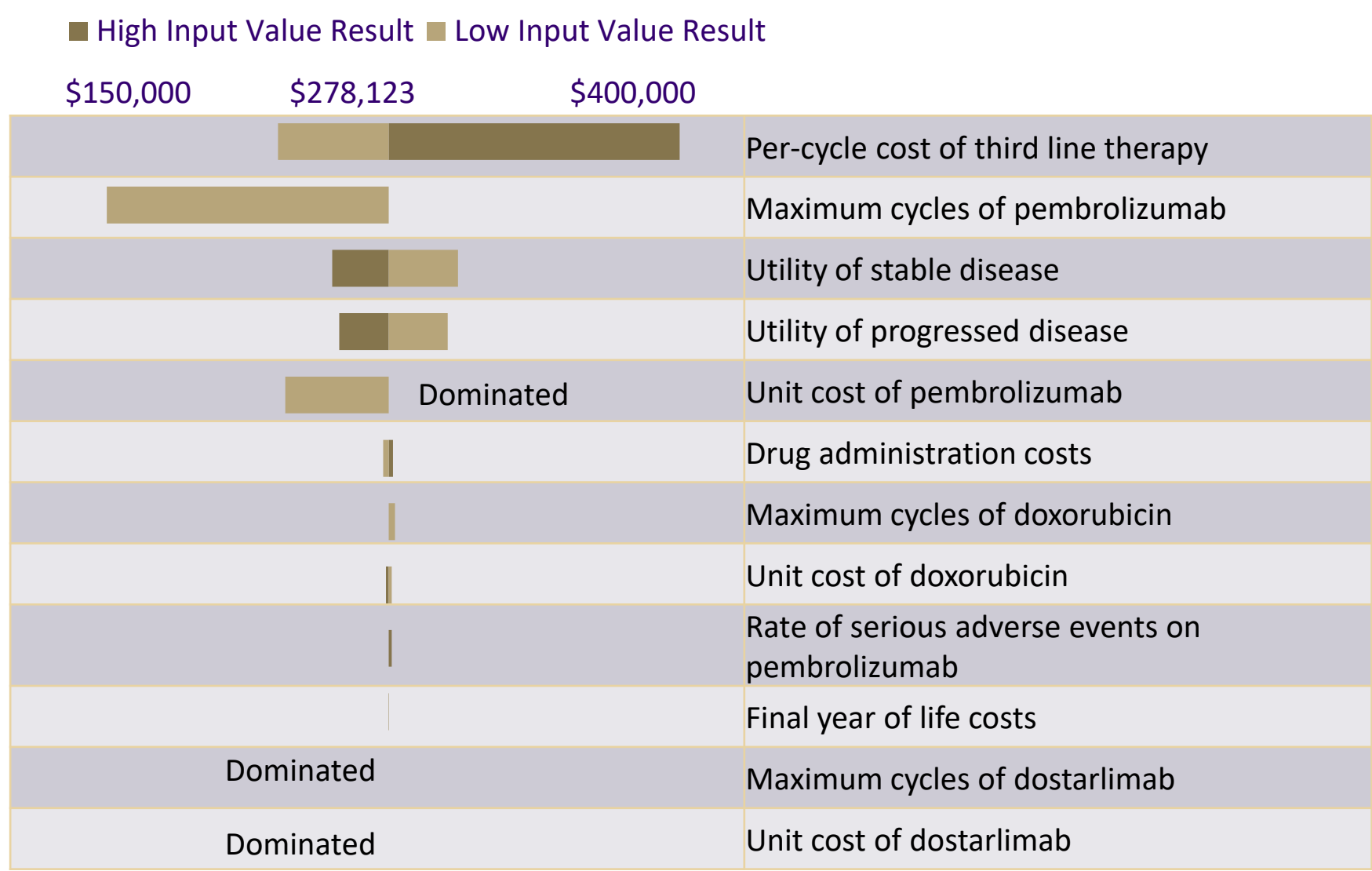


Figure 1B: One-Way Sensitivity Analysis: Dostarlimab vs PLD

