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INTRODUCTION

- In 2024, it was estimated that 11,700 Canadians will be diagnosed with non-Hodgkin lymphoma, with 5% to 10% of these cases representing are mantle cell lymphoma (MCL).^{1,2}
- Relapsed/refractory MCL (R/R-MCL) represents a particularly challenging subset of MCL, characterized by either the recurrence of the disease after initial remission or the persistence of the disease despite treatment. This phase is often associated with a poorer prognosis and a more aggressive disease course.
- Venetoclax, a selective and orally bioavailable small-molecule inhibitor of B-cell lymphoma-2, in combination with Ibrutinib, a Bruton's tyrosine kinase inhibitor (BTKis), (V+I) was evaluated for the treatment of R/R-MCL in the SYMPATICO trial.³
- Currently, there are no Canadian healthcare resource utilization (HCRU) studies published in R/R-MCL. There is a need to determine the burden of illness associated with V+I in the treatment of R/R-MCL compared to current standard of care and future anticipated treatments.

OBJECTIVES

• This study aims to estimate HCRU costs of V+I compared to current treatment options used for R/R-MCL, to inform health technology assessment agencies, institutional decision makers and healthcare professionals, from a Canadian and Quebec perspective.

METHODS

- A costing analysis was developed comparing V+I to current treatment options in Canadian and Quebec patients with R/R-MCL.
- Comparators: divided by line of treatment for R/R-MCL
- Second-line (2L): BTKis and chemo-immunotherapy (CIT)*
- Third-line (3L): brexucabtagene autoleucel (brexu-cel), venetoclax, lenalidomide and allogeneic stem cell transplant (allo-SCT) in addition to BTKis.
- Time horizon: treatment-specific median durations.
- Perspective/costs: Canadian and Quebec perspectives
- Healthcare system: direct medical costs (pretreatment, administration/monitoring and adverse event)
- Societal: direct and indirect costs (i.e., patient and caregiver lost productivity and out-of-pocket costs)
- Acquisition costs of active treatments were excluded.
- Model inputs were retrieved from product labels and were validated by Canadian clinical experts to reflect practice.

*Note: CITs are rarely used and are reserved for a small proportion of patients who are BTKi-resistant.

RESULTS

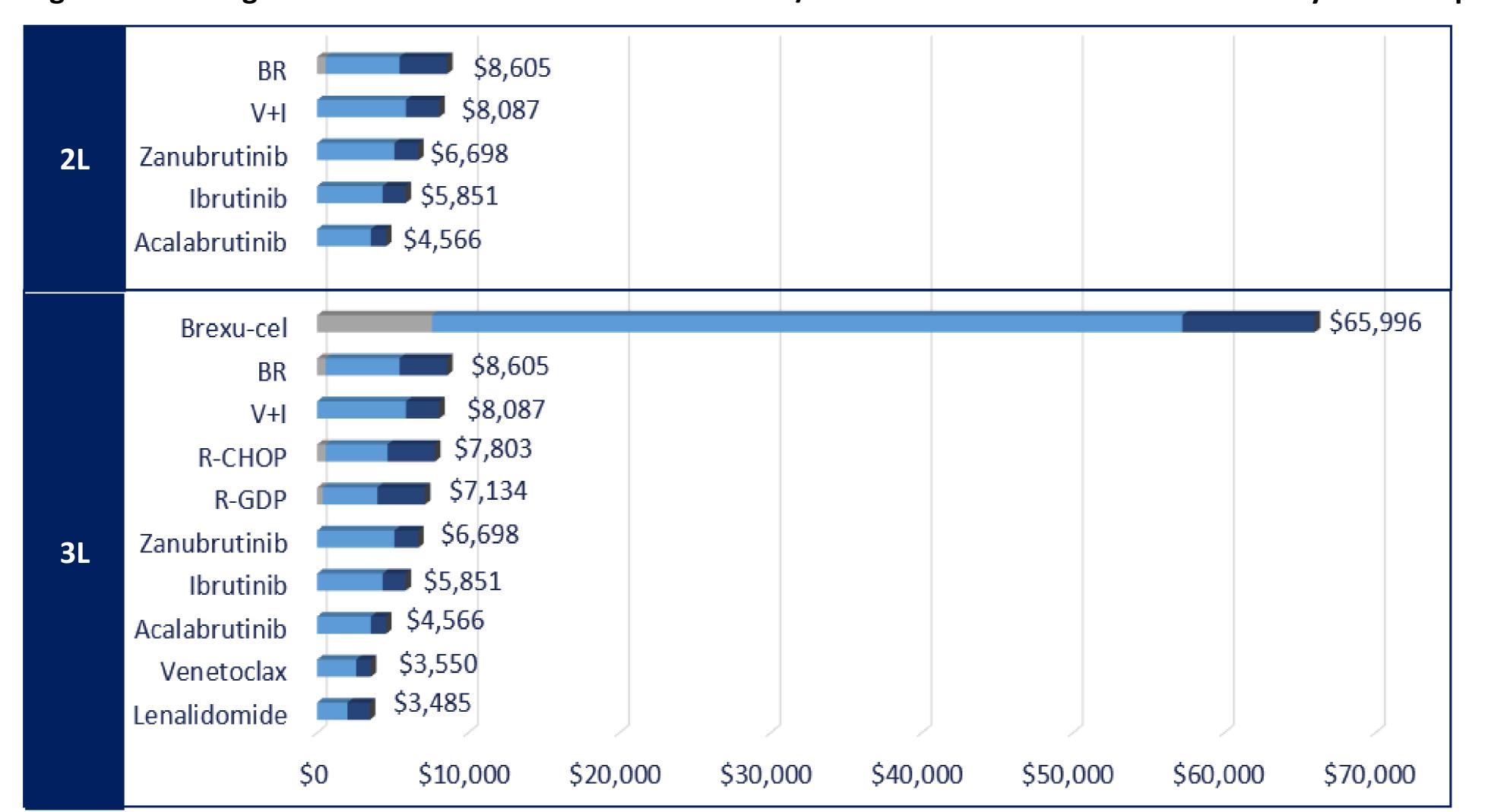
- Compared to ibrutinib, the only reimbursed BTKi in 2L, the addition of venetoclax to ibrutinib as a combination therapy generates additional HCRU cost of \$2,236 in Canada and \$1,929 in Quebec, while providing a longer median progression-free survival (PFS), as demonstrated in the SYMPATICO trial (V+I: 31.9 months vs ibrutinib: 22.1 months).
- Among 2L treatments, V+I offers HCRU cost savings compared to CITs (Table 1 & Figure 1).
- The highest HCRU cost burden in managing R/R-MCL is sourced from patients receiving 3L therapies.
- Among all treatments in 3L, brexu-cel and allo-SCT lead to the highest HCRU costs (Table 1 & Figure 1).
- Results remain similar for the Quebec setting and from a societal perspective.

Table 1. Total HCRU Costs of Treatments for R/R-MCL in Canada

Treatments	Pre- medications	Administration	Adverse Events	Productivity Loss	Healthcare system		Societal	
					Total Costs	Difference vs. V+I	Total Costs	Difference vs. V+I
				2L+				
V+I	\$0	\$5,880	\$2,207	\$5,639	\$8,087	-	\$13,726	-
Ibrutinib	\$0	\$4,342	\$1,509	\$3,144	\$5,851	\$2,236	\$8,995	\$4,732
Zanubrutinib	\$0	\$5,117	\$1,581	\$4,116	\$6,698	\$1,389	\$10,814	\$2,912
Acalabrutinib	\$0	\$3,567	\$1,000	\$2,468	\$4,566	\$3,521	\$7,034	\$6,692
BR	\$576	\$4,885	\$3,145	\$11,809	\$8,605	-\$519	\$20,414	-\$6,688
				3L+				
Venetoclax	\$0	\$2,597	\$953	\$3,133	\$3,550	\$4,537	\$6,683	\$7,043
R-CHOP	\$576	\$4,082	\$3,145	\$8,389	\$7,803	\$284	\$16,192	-\$2,465
R-GDP	\$384	\$3,605	\$3,145	\$6,768	\$7,134	\$953	\$13,902	-\$176
Brexu-cel	\$7,597	\$49,643	\$8,756	\$16,544	\$65,996	-\$57,909	\$82,540	-\$68,814
Lenalidomide	\$0	\$2,016	\$1,469	\$820	\$3,485	\$4,602	\$4,305	\$9,421
Allo-SCT	\$18,830	\$91,522	\$26,628	\$34,148	\$136,980	-\$128,893	\$171,128	-\$157,402

Abbreviations: Allo-SCT: allogeneic stem cell transplant; **BR:** bendamustine and rituximab; **Brexu-cel:** brexucabtogene autoleucel; **R-CHOP:** rituximab, cyclophosphamide, doxorubicin, vincristine and prednisone; **R-GDP:** rituximab, gemcitabine, dexamethasone and cisplatin; **V+I:** venetoclax and ibrutinib.

Figure 1. Ranking of Total HCRU Costs of Treatments for R/R-MCL in Canada from a Healthcare System Perspective



Abbreviations: Allo-SCT: allogeneic stem cell transplant; **BR:** bendamustine and rituximab; **Brexu-cel:** brexucabtogene autoleucel; **R-CHOP:** rituximab, cyclophosphamide, doxorubicin, vincristine and prednisone; **R-GDP:** rituximab, gemcitabine, dexamethasone and cisplatin; **V+I:** venetoclax and ibrutinib; **2L:** second-line; **3L:** third-line. **Note:** Allo-SCT was not included in this figure since total estimated costs are much higher than other comparators (estimated total costs of \$136,980) and would impact the visual presentation of the results within the figure.

DISCUSSION

Study Strengths:

- First study to estimate HCRU costs of all treatment options for R/R-MCL in Canada and Quebec.
- Use of a thorough validation process, involving clinical experts based in major Canadian and Quebec provinces.

Study Limits:

- Cost data primarily sourced from Ontario for the Canadian perspective.
- Study does not account for patient differences; therefore, results might not be representative for some patient subgroups of R/R-MCL.
- Although randomized controlled trials are a type of best supportive care that is often used for the treatment of R/R-MCL, they were excluded as a potential comparator since it is impossible to cost.

CONCLUSIONS

By offering a longer PFS in 2L, V+I could delay the need for a 3L treatment along with their substantial HCRU costs and burden.

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DISCLOSURES

- Jean Lachaine is a partner at PeriPharm Inc., a company that has served as a consultant to AbbVie and has received funding from AbbVie to conduct the study.
- Kimberly Guinan and Monika Ham are employees of PeriPharm Inc.
- Jean Lachaine, Kimberly Guinan and Monika Ham from PeriPharm Inc., have participated in the study conduct, data interpretation and the preparation of this abstract.
- Dr. Christopher Lemieux received honorarium from AbbVie and Jazz Pharma.
- Dr. Peter Anglin received honorarium from AbbVie, Janssen, Astrazeneca, FORUS, Sanofi, Incyte and Kite.
- Nancy Paul Roc, Harpreet Singh, Hardeek Patel, Beenish S. Manzoor and Stephane Barakat are employees of AbbVie. Employees of AbbVie may hold equity.
- AbbVie participated in the design and provided financial support for the study. AbbVie reviewed and approved this publication.
- No honoraria or payments were made for authorship.

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