# **Real-World Adherence and Healthcare Resource Utilization of Bruton Tyrosine Kinase Inhibitors in Mantle Cell Lymphoma**

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## INTRODUCTION

- Mantle cell lymphoma (MCL) is a rare and incurable B-cell malignancy<sup>1</sup>
- MCL can be controlled for a prolonged period of time but eventually becomes refractory or relapsed (R/R), leading to subsequent treatment<sup>2</sup>
- $\bullet$  Bruton tyrosine kinase inhibitors (BTKis) have been shown to be effective for the treatment of R/R MCL  $^{\rm 3-5}$
- However, real-world data for the treatment patterns and outcomes of BTKis in MCL are limited

## OBJECTIVE

 To examine treatment adherence and healthcare resource utilization (HCRU) of patients undergoing BTKi treatment in MCL

### METHODS

#### Data Source

- Data were sourced from Symphony Integrated Dataverse (IDV<sup>®</sup>, a comprehensive, longitudinal, open claims database, and integrated electronic medical record data)
- The study period was from December 2019 to November 2023 with an index period between January 2020 and October 2023

#### **Inclusion Criteria**

- Patients aged ≥18 years with ≥1 diagnosis for MCL
- Patients who initiated their first BTKi treatment or switched from ibrutinib to acalabrutinib or zanubrutinib during the index period, defined as the date of treatment initiation
- Patients with continuous enrollment in the database for 30 days prior to and 30 days after the index date

#### Cohorts

- Two cohorts were identified
- Patients receiving their first BTKi (acalabrutinib, ibrutinib, zanubrutinib) during the index period
- Patients that received ibrutinib as their first BTKi and initiated acalabrutinib or zanubrutinib as their second BTKi during the index period

#### **Study Measures**

- Demographics, clinical characteristics, and comorbidities were measured at index date
- Adherence was evaluated by compliance and persistence
- Compliance was calculated as the proportion of days covered using 30-day intervals from initiation on treatment to 1 year. Proportion of days covered > 0.80 indicated compliance
- Persistence was measured as the proportion of patients who remained on treatment among patients with sufficient follow-up periods
- HCRU was measured by all-cause outpatient visits, inpatient services, and other medical/hospital services per-patient-per-month during BTKi treatment

## RESULTS

#### First-Ever BTKi Patients

- A total of 2122 patients initiated their first-ever BTKi during the index period.
- Among these patients, 725 initiated ibrutinib, 878 initiated acalabrutinib, and 519 initiated zanubrutinib
- There were significant differences for mean age at index (P<.001), payer type (P<.001), and prior line of therapy (P<.001) (Table 1)</li>

#### **Table 1. Demographics and Clinical Characteristics**

Variable	lbrutinib (n=725)	Acalabrutinib (n=878)	Zanubrutinib (n=519)	<i>P</i> Value
Age at index				<.0001
Mean (SD)	68.31 (9.24)	70.46 (8.2)	70.04 (8.6)	
Median (IQR)	70 (62, 77)	72 (65, 77)	72 (65, 77)	
Age groups, n (%)				.0013
18-55	70 (9.66)	53 (6.04)	32 (6.17)	
56-64	167 (23.03)	161 (18.34)	96 (18.5)	
65+	488 (67.31)	664 (75.63)	391 (75.34)	
Sex				.0895
Male	542 (74.76)	613 (69.82)	375 (72.25)	
Female	183 (25.24)	265 (30.18)	144 (27.75)	
Payer type				<.0001
Medicaid	38 (5.24)	50 (5.69)	22 (4.24)	
Medicare	354 (48.83)	518 (59)	249 (47.98)	
Commercial	303 (41.79)	299 (34.05)	243 (46.82)	
Self	16 (2.21)	9 (1.03)	1 (0.19)	
Other	14 (1.93)	2 (0.23)	4 (0.77)	
Prior line of therapy	<.0001			
Mean (SD)	0.78 (0.63)	0.94 (0.61)	0.91 (0.64)	
Median (IQR)	1 (O, 1)	1 (1, 1)	1 (1, 1)	

SD, standard deviation; IQR, interquartile range.

#### Comorbidities

 Across the measured comorbidities at baseline, there were significant differences among the three BTKis for in the rate of atrial fibrillation (acalabrutinib=1.94%; ibrutinib=1.24%; zanubrutinib=3.47%, P=.0232)

#### **Compliance and Persistence**

• Compliance and persistence results are shown in **Figure 1** 

Figure 1. Compliance and Persistence in First-Ever Bruton Tyrosine Kinase Inhibitor Users



- Compliance at 1 year was numerically highest for zanubrutinib (16.73%) followed by acalabrutinib (16.45%) and ibrutinib (11.80%) (P=.1476)
- Treatment persistence at 1 and 2 years was also numerically highest for zanubrutinib (33.1%; 18.6%) compared to acalabrutinib (32.8%; 16.3%) and ibrutinib (30.6%; 15.7%) (*P*=.2180; *P*=.2275)

#### Acalabrutinib and Zanubrutinib as the Second BTKi After Ibrutinib

- A total of 228 patients switched from ibrutinib to acalabrutinib (n=140) or zanubrutinib (n=88).
- Over 85% of patients switched directly from ibrutinib to acalabrutinib or zanubrutinib
- There were no significant differences in mean age, sex, or prior lines of therapy between patients receiving acalabrutinib or zanubrutinib post-ibrutinib (Table 2)
- There were no significant differences in baseline comorbidities between patients receiving acalabrutinib or zanubrutinib post-ibrutinib

Variable	Acalabrutinib (n=140)	Zanubrutinib (n=88)	<i>P</i> Value
Age at index	.0781		
Mean (SD)	70.77 (8.7)	68.91 (7.05)	
Median (IQR)	73 (66.5, 77)	68 (64, 76)	
Age groups, n (%)	.0091		
18-55	10 (7.14)	3 (3.41)	
56-64	19 (13.57)	26 (29.55)	
65+	111 (79.29)	59 (67.05)	
Sex	.1836		
Male	118 (84.29)	68 (77.27)	
Female	22 (15.71)	20 (22.73)	
Payer type	.0201		
Medicaid	3 (2.14)	5 (5.68)	
Medicare	90 (64.29)	39 (44.32)	
Commercial	46 (32.86)	42 (47.73)	
Self	O (O)	O (O)	
Other	1 (0.71)	2 (2.27)	
Prior line of therapy	.9962		
Mean (SD)	1.36 (1)	1.36 (1.03)	
Median (IQR)	1 (1, 2)	1 (1, 2)	
Switching Status	.8654		
Direct switch from ibrutinib	122 (87.14)	76 (86.36)	

**Table 2. Demographics and Clinical Characteristics of Patients Post-Ibrutinib** 

SD, standard deviation; IQR, interquartile range.

Among patients who switched from ibrutinib to zanubrutinib or acalabrutinib

- Zanubrutinib had numerically better 1-year compliance (P=.2176)
- Zanubrutinib had numerically better treatment persistence at 1 and 2 years (P=.2687; P=.6270) (Figure 2)

#### Figure 2. Compliance and Persistence of Patients Post-Ibrutinib



## CONCLUSIONS

- In this study, zanubrutinib was associated with a trend towards improved compliance, persistence, and HCRU when used as the first BTKi and after prior ibrutinib
- Among patients who switched from ibrutinib to zanubrutinib or acalabrutinib
- Zanubrutinib demonstrated numerically better compliance and persistence than acalabrutinib
- Zanubrutinib demonstrated numerically lower outpatient visits and inpatient services than acalabrutinib
- HCRU showed that the mean (SD) number of outpatient visits and inpatient services were numerically lower in patients that switched from ibrutinib to zanubrutinib (1.12 [1.67] vs 1.62 [3.17]; P=.1755) than in patients that switched to acalabrutinib (0.22 [0.68] vs 0.68 [3.12]; P=.1693; **Figure 3**)

#### Figure 3. Healthcare Resource Utilization of Patients Post-Ibrutinib



#### REFERENCES

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#### DISCLOSURES

BS: Employment: Moffitt Cancer Center. Research Funding: Jazz Pharmaceuticals, Servier, Kite. Travel accommodations for Kite. Other relationships with DSMB, Pepromene Bio. WF: Employment and equity holder in Real Chemistry.
KE: Employment in Real Chemistry. PC: Employment in Real Chemistry.
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