ENDOCRINOPATHIES ASSOCIATED WITH IBRUTINIB – A RETROSPECTIVE ANALYSIS

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INTRODUCTION

- > Tyrosine Kinase Inhibitors (TKI) are used to for the targeted treatment of hematologic malignancies
- > FDA approved indications for ibrutinib have been steadily increasing since 2013
- > Some TKIs have been shown to have endocrine side effects (S/E)- however there is a paucity of literature regarding ibrutinib specifically

METHODOLOGY

Retrospective observational single center study of all adults who initiated ibrutinib from Nov 2013 to Jan 2020 for treatment of any malignancy or graft vs. host disease

OUTCOMES

- > **Primary Outcome:** To determine the frequency of endocrine related S/E associated with ibrutinib
- > Secondary Outcomes: To evaluate if clinical outcomes differ among patients with pre-existing endocrine comorbidities who received ibrutinib

Total patients	N = 105
Female	18%
Age	72 ± 11 years
Stage IV cancer	39%
Prior endocrinopathy	72%

> Three patients (2.8%) developed endocrine related S/E after ibrutinib initiation (Table 1)

Endocrine related side effect

Hyperparathyroidism

Hypothyroidism

Gynaecomastia

- to ibrutinib
 - Most common non endocrine S/E Gastrointestinal (N/V/diarrhea)
- Time to discontinuation of ibrutinib 10.9 months
- \blacktriangleright Most common indication for use CLL (74%)
- > No statistically significant difference in mortality between those with and without endocrine comorbidities on ibrutinib (p=0.81) (Figure 1)
- Small sample size
- Association of endocrinopathies and ibrutinib did not necessarily indicate causation



RESULTS

Age	Time from last ibrutinib dose to onset of S/E	Concurrent anti- neoplastic agents
71y	3 months	-
83y	6 months	Bendamustine Rituximab
79y	10 months	Chlorambucil

> Twenty-two patients (21%) in total developed S/E attributed

LIMITATIONS



Figure 1: Kaplan-Meier survival curves comparing all-cause mortality of patients on ibrutinib with and without endocrine co-morbidities

CONCLUSIONS

- Although many TKIs have been associated with subsequent endocrinopathies, this is not a common association with ibrutinib
- Presence of pre existing endocrine comorbidities had no effect on overall survival in patients on ibrutinib
- Although rare, early recognition of this S/E is important to facilitate prompt treatment

References

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