Danish Cancer Society

BACKGROUND & AIM

Hypertension (HTN) is highly prevalent in newly diagnosed CLL and a common adverse event in patients treated with BTK inhibitors.

Our aim was to assess the impact of HTN on treatment and survival outcomes for patients with CLL.

METHODS

We used data from nation-wide registers. Data was linked on an individual level.

- CLL cohort: The Danish CLL Register, patients diagnosed from 2008 to 2022. Last follow-up: Sep. 2022.
- Causes of death: The Danish National Register of **Causes of Death.**
- HTN diagnosis (Table 1):
 - The Danish National Patient Register and Epic
 - The Danish Prescription Register

Stratification: HTN-status prior to CLL diagnosis.

TABLE 1 – HTN definition

Two prescriptions on first choice anti-HTN or combination anti-HTN therapy, one prescribed <180 days prior to CLL diagnosis

Two prescriptions on anti-HTN of later choice, both prescribed <180 days prior to CLL diagnosis

Two ICD10 codes for HTN prior to CLL diagnosis

Primary outcome:

- Overall survival (OS) from time of CLL diagnosis and first line treatment comparing CLL patients w/ and w/o HTN.
 - Kaplan-Meier and Cox regression model.

Secondary outcomes:

- Event-free survival (EFS), time to first treatment (TTFT) and cause-specific mortality.
- Cumulative incidence and Cox regression model.

Association between hypertension and overall survival in chronic lymphocytic leukemia

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- Patient cohort: 6,557 patients, 46% of whom had HTN
- Median age: 71 years (74 vs 86 years for patients w/ and w/o HTN, respectively)
- CLL patients with HTN had a shorter overall survival (OS), particularly following first line treatment (Figure 1 and 2).
- Patients >75y had similar 5-year OS rates w/ and w/o HTN (52% vs 54%, respectively)
- For CLL patients, HTN was associated with a higher risk of CLL-unrelated and CLL-related death, and they more often died from infectious and cardiovascular causes (Figure 4).



Figure 3: OS from first line treatment with either FCR, BR or targeted drugs. stratified Figure 4: Cumulative incidence curves showing risk of death due to A: infection (HR: by treatment type for A: CLL patients w/ HTN and B: w/o HTN at time of treatment. 1.16 [95% CI 0.98;1.37] and B: hematological malignancies (HR: 1.06 [95% CI 1.86;1.31] stratified by HTN status at CLL diagnosis. Death due to other causes was treated as competing risk. Cox model adjusted for age, year of CLL diagnosis and sex. Perfomred on 288 patients who underwent medical journal review .

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Figure 1: Kaplan-Meier curve showing OS following CLL diagnosis for CLL patients w and w/o HTN

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Figure 2: Forest plot presenting Cox model hazard ratio (HR) and 95% confidence interval (CI) for OS, TTFT and EFS from CLL diagnosis and OS and EFS from first line CLL treatment comparing CLL patients w and w/o HTN (as reference). Adjusted for age at CLL diagnosis, year of CLL diagnosis, sex, CLL-IPI and numbers of comorbidities, including cardiac.

TAKE HOME MESSAGE

- HTN was associated with shorter survival following both CLL diagnosis and treatment and a longer time to treatment.
- CLL patients with HTN may be more vulnerable to infections, which could be considered upon choice of CLL treatment.
- CLL patients with HTN also benefitted from chemoimmunotherapy compared to targeted treatment.