

LETTER TO THE EDITOR

Open Access

Long lasting complete molecular remission after suspending dasatinib treatment in chronic myeloid leukemia

Klára Gadó^{1*}, András Matolcsy², Judit Csomor², Dóra Kicsi³, Csaba Bödör² and Gyula Domján¹

Abstract

Tyrosine kinase inhibitors specific for BCR-ABL, were a major breakthrough in CML therapy. Second generation tyrosine kinase inhibitors (dasatinib, nilotinib) are indicated for imatinib resistant and intolerant patients. Present guidelines recommend continuous drug dosing for maintaining remission. There is no available data concerning the optimal duration of dasatinib therapy. We report the case of an imatinib intolerant patient who succeeded a complete molecular remission with dasatinib. Dasatinib was stopped because of intolerance, but complete molecular remission was sustained for one year and minor molecular remission for 27 months after discontinuation of dasatinib.

Keywords: Chronic myeloid leukemia, Dasatinib, Imatinib, Tyrosine kinase inhibitor, Drug intolerance

Summary

Tyrosine kinase inhibitors specific for BCR-ABL, were a major breakthrough in CML therapy. Second generation tyrosine kinase inhibitors (dasatinib, nilotinib) are indicated for imatinib resistant and intolerant patients. Present guidelines recommend continuous drug dosing for maintaining remission. There is no available data concerning the optimal duration of dasatinib therapy. We report the case of an imatinib intolerant patient who succeeded a complete molecular remission with dasatinib. Dasatinib was stopped because of intolerance, but complete molecular remission was sustained for one year and minor molecular remission for 27 months after discontinuation of dasatinib.

The introduction of tyrosine kinase inhibitors specific for BCR-ABL, were a major breakthrough in CML therapy [1]. Approximately 30% of patients receiving imatinib as first-line therapy will discontinue treatment by 5 years because of imatinib resistance or drug toxicity [2].

Second generation TKIs (dasatinib, nilotinib) are indicated for those patients who are refractory or intolerant

to imatinib [3,4]. However, after stopping TKI administration, relapse comes about inevitable, so the present guidelines recommend continuous drug dosing [5]. While there is an ongoing prospective study with imatinib (Stop Imatinib study) [6], no available data arising from controlled studies exists with dasatinib concerning the optimal duration of therapy. Consequently, discontinuation of dasatinib is not recommended outside of clinical trials but remains an active area of research [7].

Our patient, a 84 year old lady with a chronic phase CML was intolerant to imatinib. She was given dasatinib as second line treatment. Though a reduced dose of 50 mg daily was applied, a complete molecular remission developed within two months. After four months she suspended taking the drug because of intolerance (pleural effusion, congestive heart failure, Adams-Stokes syndrome had been developed), but complete molecular remission was lasting till one year after stopping dasatinib. The patient did not receive any kind of anti-CML therapy during this year. After one year without any TKI, the expression level of *BCR/ABL* major fusion gene has increased gradually, but even now, 27 months after stopping dasatinib treatment, the patient is still in minor molecular remission (Figure 1).

Because of her several kind of comorbidities, her good quality of life, the actual hematological remission and

* Correspondence: gadok@freemail.hu

¹1st Department of Internal Medicine, Semmelweis University, Korányi S. Street 2, 1083, Budapest, Hungary

Full list of author information is available at the end of the article

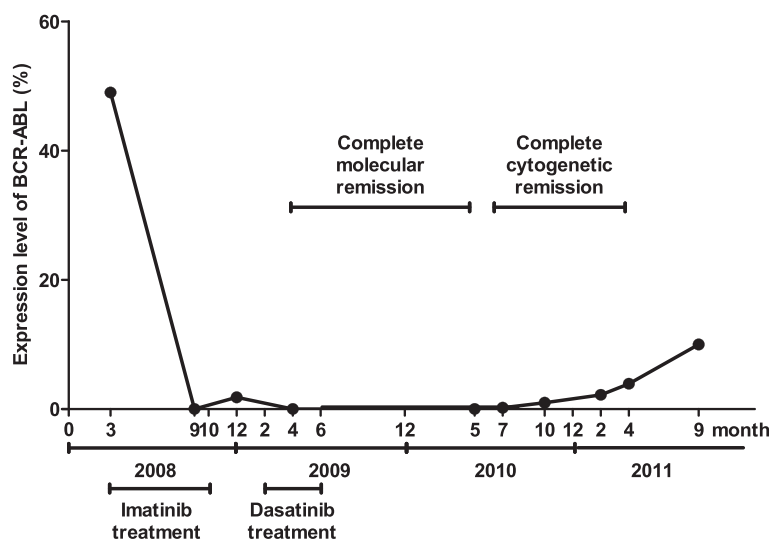


Figure 1 Expression level of BCR-ABL major fusion gene at various time during therapy measured by real-time quantitative PCR using ABL gene amplification as a control.

also her reluctance to take any kind of TKIs, our therapeutic strategy is only watch and wait.

This is the first report about maintaining complete molecular remission after one year of suspending dasatinib treatment.

Abbreviations

CML, Chronic myeloid leukemia; TKI, Tyrosine kinase inhibitor.

Competing interests

The authors have no relevant conflict of interests.

Authors' contributions

KG, DK and GD were responsible for the clinical guidance of the patient, Judit Csomor, András Matolcsy and Csaba Bődör carried out the molecular biological examinations. All authors participated in drafting and critically revising the manuscript. All authors read and approved the final manuscript.

Author details

¹1st Department of Internal Medicine, Semmelweis University, Korányi S. Street 2, 1083, Budapest, Hungary. ²1st Department of Pathology and Experimental Cancer Research, Semmelweis University, Budapest, Hungary. ³Department of Internal Medicine, St Rokus Hospital, Budapest, Hungary.

Received: 24 March 2012 Accepted: 27 April 2012

Published: 11 July 2012

References

- O'Brien SG, Guilhot F, Larson RA, Gathmann I, Baccarani M, Cervantes F, Cornelissen JJ, Fischer T, Hochhaus A, Hughes T, Lechner K, Nielsen JL, Rousselot P, Reiffers J, Saglio G, Shepherd J, Simonsson B, Gratwohl A, Goldman JM, Kantarjian H, Taylor K, Verhoef G, Bolton AE, Capdeville R, Druker BJ: **Imatinib compared with interferon and low-dose cytarabine for newly diagnosed chronic-phase chronic myeloid leukemia.** *N Engl J Med.* 2003, **348**(11):994–1004.
- Bumbea H, Vladareanu AM, Voican I, Cisleanu D, Barsan L, Onisai M: **Chronic myeloid leukemia therapy in the era of tyrosine kinase inhibitors—the first molecular targeted treatment.** *J Med Life.* 2010, **3**(2):162–6.
- Aguilera DG, Tsimberidou AM: **Dasatinib in chronic myeloid leukemia: a review.** *Ther Clin Risk Manag.* 2009, **5**(2):281–9.

- Stein B, Smith BD: **Treatment options for patients with chronic myeloid leukemia who are resistant to or unable to tolerate imatinib.** *Clin Ther.* 2010, **32**(5):804–20.
- Traer E, Deininger MW: **How much and how long: tyrosine kinase inhibitor therapy in chronic myeloid leukemia.** *Clin Lymphoma Myeloma Leuk.* 2010, **10**(Suppl 1):S20–6.
- Mahon FX, Rea D, Guilhot J, Guilhot F, Huguet F, Nicolini F, Legros L, Charbonnier A, Guerci A, Varet B, Etienne G, Reiffers J, Rousselot P: **Discontinuation of imatinib in patients with chronic myeloid leukaemia who have maintained complete molecular remission for at least 2 years: the prospective, multicentre Stop Imatinib (STIM) trial.** *Lancet Oncol.* 2010, **11**(11):1029–35.
- Baccarani M, Cortes J, Pane F, Niederwieser D, Saglio G, Apperley J, Cervantes F, Deininger M, Gratwohl A, Guilhot F, Hochhaus A, Horowitz M, Hughes T, Kantarjian H, Larson R, Radich J, Simonsson B, Silver RT, Goldman J, Hehlmann R: **Chronic myeloid leukemia: an update of concepts and management recommendations of European LeukemiaNet.** *J Clin Oncol.* 2009, **27**(35):6041–51.

doi:10.1186/2162-3619-1-17

Cite this article as: Gadó et al.: Long lasting complete molecular remission after suspending dasatinib treatment in chronic myeloid leukemia. *Experimental Hematology & Oncology* 2012 1:17.

Submit your next manuscript to BioMed Central and take full advantage of:

- Convenient online submission
- Thorough peer review
- No space constraints or color figure charges
- Immediate publication on acceptance
- Inclusion in PubMed, CAS, Scopus and Google Scholar
- Research which is freely available for redistribution

Submit your manuscript at
www.biomedcentral.com/submit

