

Title Page

- Title.

Secondary loss of response to infliximab in pediatric Crohn's disease: Does it matter how and when we start?

- Authors.

Rishi Bolia MD DM, Jeremy Rosenbaum FRACP, Vered Schildkraut FRACP, Winita Hardikar FRACP PhD, Mark Oliver FRACP MD, Donald Cameron FRACP, George Alex FRACP PhD

Department of Gastroenterology and Clinical Nutrition, The Royal Children's Hospital, Melbourne, Parkville, Australia

- Short title.

LOSS OF RESPONSE TO INFLIXIMAB IN PEDIATRIC CROHNS DISEASE

- Address for correspondence.

Dr. George Alex

Department of Gastroenterology

Royal Children's Hospital Melbourne

Ph. 613 93455060

Fax 613 93456240

E-mail – george.alex@rch.org.au

- Nothing to disclose
- Funding – none
- Word count (Excluding references) – 2575
- Tables - 1

Abstract

Objectives: A significant proportion of children with Crohn's disease develop a secondary loss of response (LOR) to infliximab. Our aim was to study the impact of initial treatment strategies on secondary LOR.

Methods: We reviewed the medical records of children with Crohn's disease who received scheduled maintenance infliximab therapy for at least 12 months. We compared children who developed LOR with those who did not; with regards to their clinical and laboratory parameters, disease phenotype and treatment strategy before developing LOR.

Results: 73 children (Median age at diagnosis- 11 (2 – 16) years, 41 boys) who had received a median duration of 33 (13-110) months of infliximab therapy were included in the final analysis. LOR was seen in 25(34.2%). Demographic variables, disease phenotype (age, disease location and behavior), inflammatory parameters and PCDAI at induction with infliximab, was similar between both groups.

Children with LOR had a significantly greater number of flares of the disease when compared to those who did not have LOR [4 (1 – 8) vs. 2(1- 5) $p = 0.03$]. The choice of the concomitant immunomodulator - Methotrexate [11/29(37.9%)] vs. Azathioprine [11/36(30.5%)] ($p = 0.6$) did not affect LOR rates. The median time-lag between diagnosis and induction with infliximab was significantly longer in children with LOR as compared to those who did not have a LOR[28 (4-90) months vs. 12.5(1 – 121) months, $p=0.004$].

Conclusion: Early use of infliximab in pediatric Crohn's disease is associated with a decrease in secondary LOR. The type of concomitant immunomodulator used does not make a difference to LOR rates.

Key-words –Infliximab, Pediatric Crohn's disease

ACCEPTED