A Comparison Between Time to Disease Activity Remission and Improvement in Patient Reported Outcomes in Rheumatoid Arthritis Patients: Results From The Ontario Best Practices Research Initiative (OBRI)

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BACKGROUND

• It is unclear if patient reported outcomes (PROs) such as fatigue and pain lag behind clinical remission in patients with RA.

OBJECTIVES

We aimed to contrast the timing of low disease state and remission with improvement in PROs in the Ontario Best Practices Research Initiative (OBRI); a clinical registry for RA (OBRI-RA registry) followed in routine care in Ontario-Canada (www.obri.ca).

METHODS

- RA patients enrolled in the OBRI were included if they had their first physician visit and first interview within 60 days' gap, and at least two visits including baseline visit, and 6 months' follow-up.
- We excluded those patients with missing data on clinical disease activity index (CDAI), swollen joint count (SJC-28), patient global assessment (PtGA), pain, or fatigue score at baseline. Patients were also excluded if they were in low/remission state based on these outcomes at baseline.
- Outcome Definition:
 - Remission: CDAI ≤ 2.8; SJC-28 ≤1.0; PtGA ≤ 1.0; pain ≤ 1.0; fatigue ≤ 1.0
 - Low disease state: CDAI ≤ 10; SJC-28 ≤2.0; PtGA ≤ 2.0; pain ≤ 2.0;
 fatigue ≤ 2.0
- Kaplan-Meier survival analysis was used to assess
 - The time to first remission based on CDAI, SJC-28, PtGA, pain, and fatigue score.
 - The time to first LDA based on CDAI, SJC-28, PtGA, pain, and fatigue score.

RESULTS

- A total of 986 patients were included. Mean (SD) age and disease duration were 57.4 (12.9) years and 8.3 (9.9) years, respectively, and the majority were females (80.0%) and established RA (65.0%)(Table 1).
- Mean (SD) of CDAI, SJC-28, PtGA, pain, and fatigue at enrolment was 29.8 (11.7), 8.3 (4.6), 6.4 (1.9), 6.6 (1.9), and 6.7 (2.0), respectively.

- During the study period CDAI remission was reported for 392 (39.8%) patients and CDAI LDA for 815 (87.2%).
- The median time to first remission and LDA was the lowest for SJC-28 (12.5 months and 9.0, respectively) among all disease measures assessed by physician and PROs outcomes (Figure 1 and Table 2).
- Median time to first pain remission was higher than time to CDAI remission (54.7 vs. 46.5 months)(Figure 1A).
- Median time to first LDA for all PROs was higher than time to first LDA based on CDAI and SJC-28 (Table 2 and Figure 1B).

Table 1: Patients Profile at enrolment

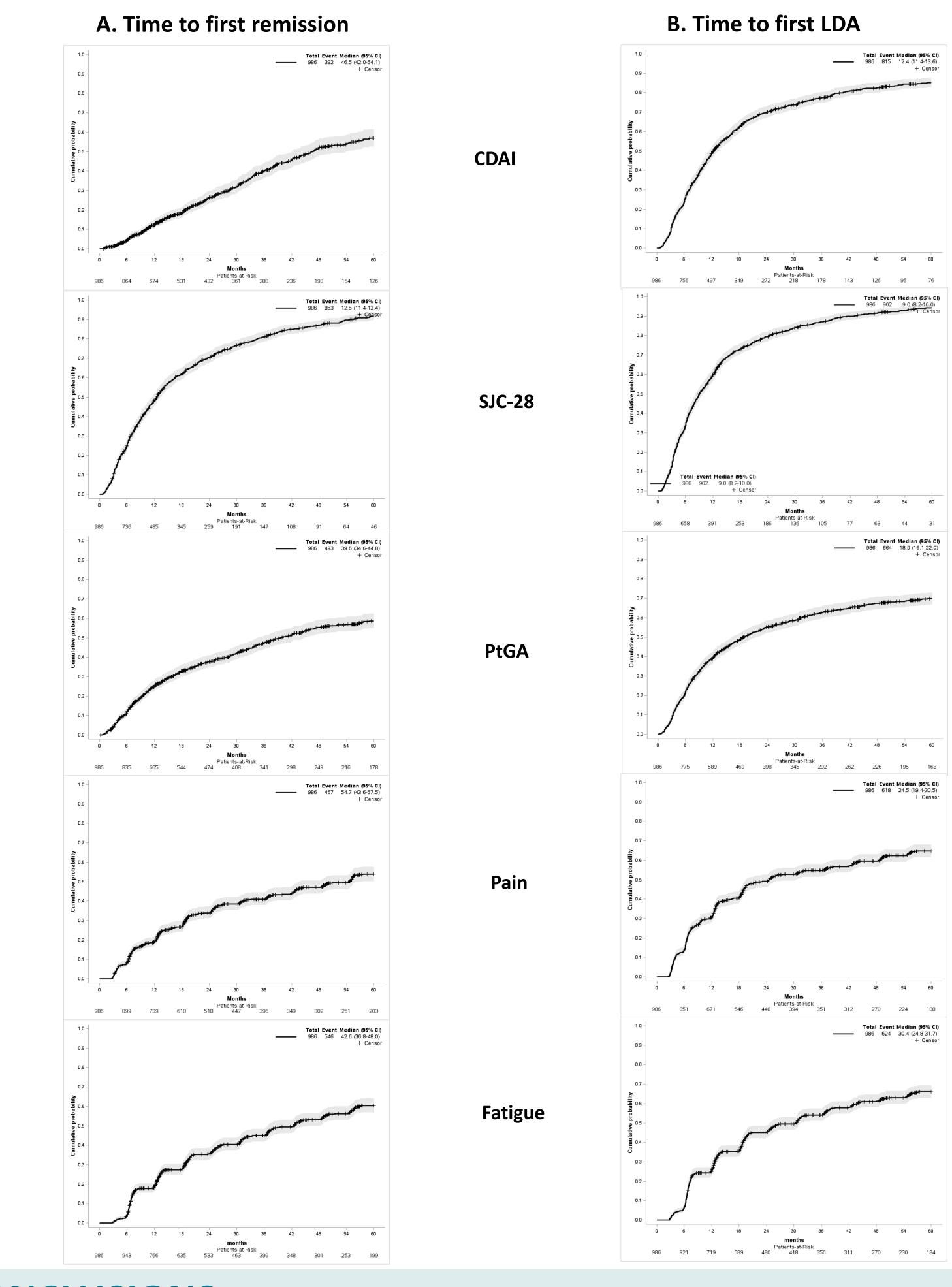
	TOTAL	
	(N=986)	
emographic Factors		
Age, years, mean (SD)	57.4 (12.9)	
Sex, Female, n (%)	789 (80.0)	
Marital status, married , n (%)	674 (68.4)	
Education status, post-secondary, n (%)	537 (54.5)	
Household annual income, >50,000CAD, n (%) (available = 781)	428 (43.4)	
Health insurance coverage, (OHIP +private or ODB), n (%)	831 (84.3)	
Residential area, Urban, n (%)	796 (80.7)	
Smoking history, n (%)		
Never smoked	437 (44.3)	
Former smoker	357 (36.2)	
Current smoker	192 (19.5)	
Disease Factors		
Disease duration since diagnosis, years, mean (SD)	8.3 (9.9)	
Early RA (≤ 1 year), n (%)	347 (35.0)	
RF positive, n (%) (available = 926)	658 (66.7)	
Presence of erosion, n (%)	400 (40.6)	
SJC-28 (0-28), mean (SD)	8.3 (4.6)	
TJC-28 (0-28), mean (SD)	9.3 (6.6)	
PhGA (0-10), mean (SD) (available =919)	5.7 (2.0)	
PtGA (0-10), mean (SD)	6.4 (1.9)	
ESR, mean (SD) (n=867)	26.9 (22.9)	
CRP, mean (SD) (n=798)	15.4 (22.8)	
CDAI (0-76), mean (SD)	29.8 (11.7)	
HAQ-DI (0-3), mean (SD)	1.6 (0.62)	
Pain (0-10), mean (SD)	6.6 (1.9)	
Fatigue (0-10), mean (SD)	6.7 (2.0)	
omorbidities		
Number of comorbidities, mean (SD)	3.8 (2.6)	
CVD, n (%)	142 (11.5)	
Hypertension, n (%)	376 (38.1)	
Diabetes Mellitus, n (%)	113 (15.6)	
Lung diseases, n (%)	154 (15.6)	
Nedication Factors		
Prior use of bDMARDs, n (%)	285 (28.9)	
Prior use of csDMARDs, n (%)	789 (80.0)	
Current bDMARDs use, n (%)	196 (19.9)	
Current csDMARDs use, n (%)	877 (89.0)	
Number of csDMARDs, mean (SD)	1.45 (0.85)	
Current steroid use, n (%)	216 (21.9)	

OHIP: Ontario health insurance plan; ODB: Ontario drug benefit; RF: Rheumatoid factor; SJC: Swollen Joint count; TJC: Tender joint count; PhGA: Physician global assessment; PtGA: Patient global assessment; ESR: Erythrocyte sedimentation rate; CRP: C-reactive Protein; CDAI: Clinical disease activity index; HAQ-DI: Health assessment questionnaire disability index; CVD: Cardiovascular disease; bDMARDs: biologic disease modifying antirheumatic drugs; csDMARDs: conventional synthetic disease modifying antirheumatic drugs.

Table 2: Median Time to First Remission and LDA Based on CDAI, SJC-28, and PROs

	N=986	
	n (%)	Median survival time, months (95% CI)
First Remission		
- CDAI remission (CDAI ≤ 2.8)	392 (39.8)	46.5 (42.0-54.1)
- SJC-28 remission (SJC28 ≤ 1.0)	853 (86.5)	12.5 (11.4-13.4)
- PtGA remission (PtGA ≤ 1.0)	493 (50.0)	39.6 (34.6-44.8)
- Pain remission (Pain≤ 1.0)	467 (47.4)	54.7 (43.6-57.5)
- Fatigue remission (Fatigue ≤ 1.0)	546 (55.4)	42.6 (36.8-48.0)
First Low disease state		
- CDAI LDA (CDAI ≤ 10)	815 (82.7)	12.4 (11.4-13.6)
- SJC-28 LDA (SJC28 ≤ 2.0)	902 (91.5)	9.0 (8.2-10.0)
- PtGA LDA (PtGA ≤ 2.0)	664 (67.3)	18.9 (16.1-22.0)
- Pain LDA (Pain≤ 2.0)	618 (62.7)	24.5 (19.4-30.5)
- Fatigue LDA (Fatigue ≤ 2.0)	624 (63.3)	30.4 (24.8-31.7)

Figure 1: Time to First Remission and LDA Based on CDAI, SJC-28, and PROs



CONCLUSIONS

- Our study shows that PRO improvement, particularly pain, lag behind CDAI and SJC-28 remission/LDA in patients with RA.
- Future work will include a subset analysis to determine time to remission/LDA in those achieving CDAI/SJC-28 remission/LDA before PRO remission/LDA.

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