

Supplementary file

Adverse events of special interest (AESIs) were recorded separately and included infections (opportunistic and serious infections as defined by treatment with intravenous anti-infectives), myocardial infarction/acute coronary syndrome, gastrointestinal perforation and related events, malignant tumours, anaphylactic/hypersensitivity reactions, demyelinating diseases, stroke, bleeding, and hepatic events.

Supplementary Table S1. Reductions in disease activity and physical functioning over the course of the ICHIBAN study according to rheumatoid arthritis onset.

	Week 104			Last visit		
	Early-onset (n=403)	Late-onset (n=207)	<i>p</i> -value	Early-onset (n=403)	Late-onset (n=207)	<i>p</i> -value
DAS28-ESR, mean reduction (95% CI)	2.95 (2.66–3.23)	3.20 (2.81–3.58)	0.34	2.21 (2.03–2.39)	2.30 (2.04–2.56)	0.60
CDAI, mean reduction (95% CI)	18.69 (15.97–21.42)	19.78 (16.45–23.12)	0.67	14.79 (13.33–16.26)	13.48 (11.37–15.58)	0.31
HAQ-DI, mean reduction (95% CI)	0.28 (0.16–0.40)	0.33 (0.15–0.52)	0.64	0.20 (0.14–0.26)	0.21 (0.12–0.29)	0.95

Patients >65 years diagnosed with rheumatoid arthritis prior to 65 years of age were considered early-onset and patients diagnosed with rheumatoid arthritis after 65 years of age were considered late-onset.

CDAI: Clinical Disease Activity Index; CI: confidence interval; DAS28-ESR: Disease Activity Score-28 erythrocyte sedimentation rate; HAQ-DI: Health Assessment Questionnaire Disease Index.

Supplementary Table S2. Treatment-emergent adverse events in patients with the outcome of death in patients >65 years old at baseline.

Sex	Age	MedDRA SOC	Preferred term
Female	77	Surgical and medicinal procedures	Surgery
Female	77	General disorders and administration site conditions	Death
Male	75	Psychiatric disorders	Delirium
		Respiratory, thoracic and mediastinal disorders	Lung disorder
		Infections and infestations	Infection
Female	75	Gastrointestinal disorders	Large intestine perforation
Male	82	Infections and infestations	Pneumonia
Female	76	General disorders and administration site conditions	Death
Male	69	Injury, poisoning and procedural complications	Anaemia postoperative
Male	70	Cardiac disorders	Electromechanical dissociation
		Cardiac disorders	Arrhythmia
		Cardiac disorders	Acute myocardial infarction
Female	74	Cardiac disorders	Myocardial infarction
Female	68	Infections and infestations	Empyema
		Respiratory, thoracic and mediastinal disorders	Pulmonary embolism
Male	67	Cardiac disorders	Myocardial infarction
Male	67	General disorders and administration site conditions	Death
Female	69	Infections and infestations	Pneumonia
		Renal and urinary disorders	Renal failure
Female	67	Hepatobiliary disorders	Gallbladder perforation

MedDRA: Medical Dictionary for Regulatory Activities; SOC: system organ class.