

Additional file 4. Common, all-causality adverse events reported in patients with OS ≥ 24 versus < 24 months in the placebo titration arm

Adverse event, ^a n (%)	OS ≥ 24 months, n = 30		OS < 24 months, n = 20 ^b	
	All Grades	Grade ≥ 3	All Grades	Grade ≥ 3
Any	30 (100)	14 (47)	17 (85)	12 (60)
Diarrhoea	24 (80)	3 (10)	8 (40)	0
Hypertension	15 (50)	3 (10)	7 (35)	2 (10)
Fatigue	15 (50)	2 (7)	9 (45)	0
Nausea	14 (47)	0	3 (15)	0
Hand-foot syndrome	8 (27)	1 (3)	2 (10)	0
Hypothyroidism	8 (27)	0	5 (25)	0
Decreased appetite	10 (33)	0	6 (30)	0
Weight decrease	8 (27)	3 (10)	4 (20)	0
Vomiting	8 (27)	0	2 (10)	0
Dysphonia	14 (47)	0	4 (20)	0
Arthralgia	9 (30)	2 (7)	2 (10)	0
Proteinuria	9 (30)	0	2 (10)	0
Headache	11 (37)	1 (3)	1 (5)	0

Abbreviation: OS overall survival

^a Reported by $\geq 25\%$ in either group.

^b Excluded 6 patients who were censored < 24 months in OS.