

TABLE S1 Frequency of subjects experiencing TEAEs^a and number of events summarized per treatment (Trial 1)

MedDRA ^b system organ class ^c MedDRA preferred term ^c	Statistic	MHAA4549A					Placebo
		1.5 mg/kg	5 mg/kg	15 mg/kg	45 mg/kg	Overall	
Number of subjects dosed	N	4	4	4	4	16	5
Number of TEAEs	E	4	7	4	4	19	4
Number of subjects ^d with TEAEs	n (%)	2 (50)	3 (75)	2 (50)	3 (75)	10 (62.5)	3 (60)
Nervous system disorders	n (%) E	0	2 (50) 2	1 (25) 1	2 (50) 2	5 (31.3) 5	1 (20) 1
Headache	n (%) E	0	2 (50) 2	1 (25) 1	1 (25) 1	4 (25) 4	1 (20) 1
Dysgeusia	n (%) E	0	0	0	1 (25) 1	1 (6.3) 1	0
Gastrointestinal disorders	n (%) E	0	2 (50) 2	1 (25) 3	1 (25) 1	4 (25) 6	0
Abdominal pain	n (%) E	0	0	0	1 (25) 1	1 (6.3) 1	0
Abdominal pain upper	n (%) E	0	1 (25) 1	0	0	1 (6.3) 1	0
Eruption	n (%) E	0	0	1 (25) 1	0	1 (6.3) 1	0
Flatulence	n (%) E	0	0	1 (25) 1	0	1 (6.3) 1	0
Nausea	n (%) E	0	0	1 (25) 1	0	1 (6.3) 1	0
Toothache	n (%) E	0	1 (25) 1	0	0	1 (6.3) 1	0
Respiratory, thoracic and mediastinal disorders	n (%) E	1 (25) 2	1(25) 2	0	1 (25) 1	3 (18.8) 5	0
Oropharyngeal pain	n (%) E	1 (25) 1	0	0	2 (25) 1	2 (12.5) 2	0
Cough	n (%) E	0	1 (25) 1	0	0	1 (6.3) 1	0
Nasal congestion	n (%) E	1 (25) 1	0	0	0	1 (6.3) 1	0
Rhinorrhoea	n (%) E	0	1 (25) 1	0	0	1 (6.3) 1	0
Injury, poisoning and procedural complications	n (%) E	2 (50) 2	0	0	0	2 (12.5) 2	1 (20) 1
Ligament sprain	n (%) E	1 (25) 1	0	0	0	1 (6.3) 1	0
Tooth fracture	n (%) E	1 (25) 1	0	0	0	1 (6.3) 1	0
Burns second degree	n (%) E	0	0	0	0	0	1 (20) 1
Investigations	n (%) E	0	1 (25) 1	0	0	1 (6.3) 1	1 (20) 1
Alanine aminotransferase increased	n (%) E	0	1 (25) 1	0	0	1 (6.3) 1	0
Blood creatine phosphokinase increased	n (%) E	0	0	0	0	0	1 (20) 1
Skin and subcutaneous tissue disorders	n (%) E	0	0	0	0	0	1 (20) 1
Urticaria	n (%) E	0	0	0	0	0	1 (20) 1

^aTEAE, treatment-emergent adverse event^bMedDRA, Medical Dictionary for Regulatory Activities; MedDRA Version 16.0^cSystem organ class and preferred term are presented in descending order of overall incidence rate considering all active subjects.^dEach subject could only contribute once to each of the incidence rates, regardless of the number of occurrences.

TABLE S2 Frequency of subjects experiencing TEAEs^a and number of events summarized per treatment (Trial 2)

MedDRA ^b system organ class ^c MedDRA preferred term ^c	Statistic	MHAA4549A			Placebo
		8400 mg	10800 mg	Overall	
Number of subjects dosed	N	4	4	8	6
Number of TEAEs	E	15	9	24	10
Number of subjects ^d with TEAEs	n (%)	3 (75)	4 (100)	7 (87.5)	5 (83.3)
Nervous system disorders	n (%) E	3 (75) 3	1 (25) 1	4 (50) 4	2 (33.3) 4
Headache	n (%) E	3 (75) 3	1 (25) 1	4 (50) 4	2 (33.3) 4
Infections and infestations	n (%) E	1 (25) 3	2 (50) 2	3 (37.5) 5	2 (33.3) 3
Nasopharyngitis	n (%) E	1 (25) 1	2 (50) 2	3 (37.5) 3	2 (33.3) 3
Bacterial infection	n (%) E	1 (25) 2	0	1 (12.5) 2	0
General disorders and administration site conditions	n (%) E	2 (50) 2	1 (25) 1	3 (37.5) 3	0
Asthenia	n (%) E	0	1 (25) 1	1 (12.5) 1	0
Fatigue	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Peripheral swelling	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Gastrointestinal disorders	n (%) E	1 (25) 2	1 (25) 1	2 (25) 3	0
Abdominal discomfort	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Faeces soft	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Nausea	n (%) E	0	1 (25) 1	1 (12.5) 1	0
Respiratory, thoracic and mediastinal disorders	n (%) E	1 (25) 1	1 (25) 1	2 (25) 2	2 (33.3) 2
Nasal congestion	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Nasal discomfort	n (%) E	0	1 (25) 1	1 (12.5) 1	0
Oropharyngeal discomfort	n (%) E	0	0	0	1 (16.7) 1
Rhinorrhoea	n (%) E	0	0	0	1 (16.7) 1
Investigations	n (%) E	0	1 (25) 2	1 (12.5) 2	0
Blood creatine phosphokinase increased	n (%) E	0	1 (25) 2	1 (12.5) 2	0
Reproductive system and breast disorders	n (%) E	1 (25) 2	0	1 (12.5) 2	0
Metrorrhagia	n (%) E	1 (25) 2	0	1 (12.5) 2	0
Eye disorders	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Conjunctival hyperaemia	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Injury, poisoning and procedural complications	n (%) E	0	1 (25) 1	1 (12.5) 1	0
Contusion	n (%) E	0	1 (25) 1	1 (12.5) 1	0
Skin and subcutaneous tissue disorders	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Pruritus	n (%) E	1 (25) 1	0	1 (12.5) 1	0
Musculoskeletal and connective tissue disorders	n (%) E	0	0	0	1 (16.7) 1
Arthralgia	n (%) E	0	0	0	1 (16.7) 1

^aTEAE, treatment-emergent adverse event.^bMedDRA, Medical Dictionary for Regulatory Activities; MedDRA Version 17.1.^cSystem organ class and preferred term are presented in descending order of overall incidence rate considering all active subjects.^dEach subject could only contribute once to each of the incidence rates, regardless of the number of occurrences.