

Table 14.3.1.2c Summary of Adverse Events by System Organ Class, Preferred Term, Treatment and Overall (Safety Population)

Study Period: Follow-Up Period

Body System MedDRA Preferred Term	rhNGF 10 µg/ml (N=62)			rhNGF 20 µg/ml (N=65)			Vehicle Control (N=29)			Total (N=156)		
	N*	n	%	N*	n	%	N*	n	%	N*	n	%
Any Adverse Events	77	22	(35.5%)	65	21	(32.3%)	21	11	(37.9%)	163	54	(34.6%)
Eye disorders	29	14	(22.6%)	26	13	(20.0%)	12	8	(27.6%)	67	35	(22.4%)
Corneal epithelium defect	0	0		6	6	(9.2%)	0	0		6	6	(3.8%)
Blepharitis	1	1	(1.6%)	2	2	(3.1%)	2	2	(6.9%)	5	5	(3.2%)
Corneal erosion	5	2	(3.2%)	2	2	(3.1%)	0	0		7	4	(2.6%)
Cataract	3	3	(4.8%)	0	0		0	0		3	3	(1.9%)
Corneal neovascularisation	2	2	(3.2%)	3	1	(1.5%)	0	0		5	3	(1.9%)
Dry eye	1	1	(1.6%)	1	1	(1.5%)	1	1	(3.4%)	3	3	(1.9%)
Keratitis	0	0		5	3	(4.6%)	0	0		5	3	(1.9%)
Neurotrophic keratopathy	3	2	(3.2%)	1	1	(1.5%)	0	0		4	3	(1.9%)
Ulcerative keratitis	3	3	(4.8%)	0	0		0	0		3	3	(1.9%)
Conjunctival haemorrhage	2	2	(3.2%)	0	0		0	0		2	2	(1.3%)
Conjunctivitis	1	1	(1.6%)	1	1	(1.5%)	0	0		2	2	(1.3%)
Corneal opacity	1	1	(1.6%)	0	0		1	1	(3.4%)	2	2	(1.3%)
Eye pain	0	0		1	1	(1.5%)	1	1	(3.4%)	2	2	(1.3%)
Lacrimation decreased	1	1	(1.6%)	1	1	(1.5%)	0	0		2	2	(1.3%)
Ocular hyperaemia	0	0		0	0		2	2	(6.9%)	2	2	(1.3%)

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	N*	n	%	N*	n	%	N*	n	%	N*	n	%
Eye disorders												
Eyelid pain	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Eyelid ptosis	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Foreign body sensation in eyes	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Lacrimation increased	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Lenticular opacities	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Meibomianitis	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Ocular discomfort	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Ocular vascular disorder	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Pseudopterygium	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Retinal cyst	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Uveitis	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Visual acuity reduced	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Vitreous haemorrhage	2	1	(1.6%)	0	0		0	0		2	1	(0.6%)
Infections and infestations	12	8	(12.9%)	11	6	(9.2%)	3	3	(10.3%)	26	17	(10.9%)
Nasopharyngitis	3	3	(4.8%)	1	1	(1.5%)	2	2	(6.9%)	6	6	(3.8%)
Helicobacter gastritis	2	2	(3.2%)	0	0		0	0		2	2	(1.3%)

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	N*	n	%	N*	n	%	N*	n	%	N*	n	%
Infections and infestations												
Herpes ophthalmic	2	2	(3.2%)	0	0		0	0		2	2	(1.3%)
Keratitis herpetic	0	0		3	2	(3.1%)	0	0		3	2	(1.3%)
Bacterial infection	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Blister infected	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Catheter site infection	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Corneal abscess	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Corneal infection	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Cystitis	0	0		2	1	(1.5%)	0	0		2	1	(0.6%)
Herpes simplex ophthalmic	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Herpes zoster ophthalmic	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Onychomycosis	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Sinusitis	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Tonsillitis	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Urinary tract infection	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)

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	N*	n	%	N*	n	%	N*	n	%	N*	n	%
General disorders and administration site conditions	4	3	(4.8%)	5	3	(4.6%)	0	0		9	6	(3.8%)
Disease progression	2	2	(3.2%)	3	1	(1.5%)	0	0		5	3	(1.9%)
Inflammation	1	1	(1.6%)	1	1	(1.5%)	0	0		2	2	(1.3%)
Impaired healing	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Instillation site pain	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Injury, poisoning and procedural complications	3	3	(4.8%)	2	1	(1.5%)	3	2	(6.9%)	8	6	(3.8%)
Clavicle fracture	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Facial bones fracture	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Fall	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Femur fracture	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Pelvic fracture	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Tibia fracture	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Transplant failure	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Upper limb fracture	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Investigations	2	2	(3.2%)	3	3	(4.6%)	1	1	(3.4%)	6	6	(3.8%)
Intraocular pressure increased	1	1	(1.6%)	2	2	(3.1%)	1	1	(3.4%)	4	4	(2.6%)
Blood pressure increased	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)

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	N*	n	%	N*	n	%	N*	n	%	N*	n	%
Investigations												
Heart rate increased	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Gastrointestinal disorders	3	3	(4.8%)	2	2	(3.1%)	0	0		5	5	(3.2%)
Abdominal pain upper	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Dental discomfort	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Diverticular perforation	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Gastric polyps	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Nausea	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Metabolism and nutrition disorders	3	2	(3.2%)	4	3	(4.6%)	0	0		7	5	(3.2%)
Decreased appetite	2	1	(1.6%)	0	0		0	0		2	1	(0.6%)
Gout	0	0		2	1	(1.5%)	0	0		2	1	(0.6%)
Hyperuricaemia	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Hypokalaemia	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Vitamin D deficiency	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)

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	N*	n	%	N*	n	%	N*	n	%	N*	n	%
Nervous system disorders	1	1	(1.6%)	4	4	(6.2%)	0	0		5	5	(3.2%)
Brain oedema	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Burning sensation	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Headache	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Syncope	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Visual field defect	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Vascular disorders	7	5	(8.1%)	0	0		0	0		7	5	(3.2%)
Aortic dissection	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Aortic rupture	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Deep vein thrombosis	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Diabetic vascular disorder	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Diastolic hypotension	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Hypertension	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Shock haemorrhagic	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Cardiac disorders	4	4	(6.5%)	0	0		0	0		4	4	(2.6%)
Arrhythmia	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Cardiac failure	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)

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	N*	n	%	N*	n	%	N*	n	%	N*	n	%
Cardiac disorders												
Coronary artery stenosis	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Myocardial infarction	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Respiratory, thoracic and mediastinal disorders												
Respiratory failure	0	0		1	1	(1.5%)	1	1	(3.4%)	2	2	(1.3%)
Dyspnoea	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Epistaxis	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Renal and urinary disorders												
Renal colic	0	0		2	2	(3.1%)	0	0		2	2	(1.3%)
Bladder prolapse	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Immune system disorders												
Corneal graft rejection	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Immunodeficiency	0	0		2	1	(1.5%)	0	0		2	1	(0.6%)

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Neoplasms benign, malignant and unspecified (incl cysts and polyps)	0	0		1	1	(1.5%)	1	1	(3.4%)	2	2	(1.3%)
Neoplasm recurrence	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Neurofibroma	0	0		0	0		1	1	(3.4%)	1	1	(0.6%)
Blood and lymphatic system disorders	3	1	(1.6%)	0	0		0	0		3	1	(0.6%)
Anaemia	3	1	(1.6%)	0	0		0	0		3	1	(0.6%)
Hepatobiliary disorders	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Cholelithiasis	0	0		1	1	(1.5%)	0	0		1	1	(0.6%)
Musculoskeletal and connective tissue disorders	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Intervertebral disc protrusion	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Reproductive system and breast disorders	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Uterine prolapse	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Skin and subcutaneous tissue disorders	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)
Diabetic foot	1	1	(1.6%)	0	0		0	0		1	1	(0.6%)

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