

APPENDIX 2.2: Cumulative and Interval Summary Tabulation of Serious and Non-Serious Adverse Reactions from Post-Marketing Data Sources

BNT162B2

Cumulative Reporting Period: Through 18-JUN-2022

Interval Reporting Period: 19-DEC-2021 Through 18-JUN-2022

Total Number of Cases: 507,683 (Interval) / 1,485,027 (Cumulative)

Total Number of Adverse Events (PT): 1,591,026 (Interval) / 4,964,106 (Cumulative)

MedDRA Version: v.25.0J

System Organ Class

Blood and lymphatic system disorders

<i>Preferred Term</i>	<i>Total # of Spontaneous AE</i>	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Abdominal lymphadenopathy	41	9	17	17	24		
Abnormal clotting factor	13		8	1	5		
Acquired amegakaryocytic thrombocytopenia	1		1				
Acquired factor V deficiency	1	1	1				
Acquired factor VIII deficiency	2	1	2				
Acquired haemophilia	65	29	65				4
Acquired Von Willebrand's disease	3		3				
Activated protein C resistance	1		1				
Agranulocytosis	33	8	33				
Anaemia	1225	224	715	187	510	4	16
Anaemia folate deficiency	6	1	3	2	3		
Anaemia macrocytic	32	3	20	4	12		
Anaemia megaloblastic	3		2	1	1		
Anaemia neonatal						2	2
Anaemia of chronic disease	1				1		
Anaemia of pregnancy	1			1	1		
Anaemia vitamin B12 deficiency	15	4	7	3	8		
Anisocytosis	9	3	5		4		
Antiphospholipid syndrome	79	36	79				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Blood and lymphatic system disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Aplasia pure red cell	6	1	6				
Aplastic anaemia	49	18	49				
Atypical haemolytic uraemic syndrome	6	3	6				
Autoimmune anaemia	8	1	8				
Autoimmune haemolytic anaemia	129	44	129			1	1
Autoimmune neutropenia	10	3	10				
Autoimmune pancytopenia	1		1				
Bandaemia	1				1		
Basophilia	3				3		
Bicytopenia	22	5	22				
Blood disorder	116	22	55	23	61	1	1
Blood loss anaemia	32	15	32				
Bone marrow disorder	14		10	1	4		
Bone marrow failure	20	8	20				
Bone marrow infiltration	1		1				
Bone marrow oedema	29	5	12	14	17		
Bone marrow oedema syndrome	2	1	1	1	1		
Coagulation disorder neonatal	1		1				
Coagulation factor deficiency	3	1	1	1	2		
Coagulopathy	501	79	297	85	204	1	1
Cold type haemolytic anaemia	7	3	7				
Coombs negative haemolytic anaemia	6		6				
Coombs positive haemolytic anaemia	7	4	7				
Cyclic neutropenia	1			1	1		
Cytopenia	23	10	23				
Deficiency anaemia	2				2		
Dermatopathic lymphadenopathy	1				1		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Blood and lymphatic system disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Disseminated intravascular coagulation	115	31	115				
Disseminated intravascular coagulation in newborn	2		2				
Dysglobulinaemia	1				1		
Elephantiasis	4	1	4				
Elliptocytosis	2		1		1		
Eosinopenia	2				2		
Eosinophilia	154	21	89	25	65		1
Erythropenia	6	2	5	1	1		
Erythropoiesis abnormal	2	1	2				
Evans syndrome	10	1	10				
Extramedullary haemopoiesis	1				1		
Factor IX inhibition	1	1	1				
Factor VIII inhibition	2		2				
Febrile bone marrow aplasia	3	1	3				
Febrile neutropenia	26	9	26				
Foetal anaemia	1		1				
Granulocytopenia	5	1	5				
Granulocytosis	1				1		
Granulomatous lymphadenitis	2	2	2				
Haemoconcentration	2	1	1		1		
Haemoglobinaemia	3		3				
Haemolysis	73	28	73				
Haemolytic anaemia	88	29	88			2	2
Haemolytic uraemic syndrome	8	2	8				
Haemorrhagic diathesis	90	24	90				
Haemorrhagic disorder	47	13	47				
Heparin-induced thrombocytopenia	15	6	15				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Blood and lymphatic system disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Hilar lymphadenopathy	40	3	13	9	27		
Hyperchromic anaemia	4		3	1	1		
Hypercoagulation	66	13	41	12	25		
Hypereosinophilic syndrome	11	3	11				
Hyperfibrinogenaemia	10		9	1	1		
Hypergammaglobulinaemia	22	2	3	9	19		
Hyperglobulinaemia	3			1	3		
Hyperleukocytosis	152	44	152				
Hyperplasia of thymic epithelium	1				1		
Hyperprothrombinaemia	1		1				
Hypersplenism	1				1		
Hyperviscosity syndrome	1		1				
Hypochromasia	5		1		4		
Hypochromic anaemia	14	5	9	3	5		
Hypocoagulable state	3		2		1		
Hypofibrinogenaemia	2	1	1		1		
Hypoplastic anaemia	2		2				
Hypoprothrombinaemia	1		1				
Hyposplenism	1	1	1				
Idiopathic CD4 lymphocytopenia	1		1				
Idiopathic neutropenia	1		1				
Immune thrombocytopenia	893	245	893			2	6
Increased tendency to bruise	250	19	82	93	168		1
Intravascular haemolysis	2	1	2				
Iron deficiency anaemia	110	16	57	22	53	1	1
Isoimmune haemolytic disease	1		1				
Jaundice acholuric	2		2				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Blood and lymphatic system disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Leukaemoid reaction	1		1				
Leukocyte vacuolisation	1	1	1				
Leukocytosis	280	45	141	47	139		1
Leukopenia	252	77	252				1
Lupus anticoagulant hypoprothrombinaemia syndrome	1	1	1				
Lymphadenitis	2521	126	433	939	2088		2
Lymphadenocyst	1				1		
Lymphadenopathy	79171	2768	10631	28302	68540	26	80
Lymphadenopathy mediastinal	60	14	31	9	29		
Lymphatic disorder	55	8	18	14	37	1	1
Lymphatic insufficiency	3		2		1		
Lymphatic obstruction	6	2	3	1	3		
Lymph node calcification	1			1	1		
Lymph node fibrosis	2			1	2		
Lymph node haemorrhage	4	1	2		2		
Lymph node pain	9199	517	1999	2451	7200	2	19
Lymph node rupture	1				1		
Lymph node ulcer	1		1				
Lymphocytic infiltration	23	3	12	5	11		
Lymphocytosis	65	12	30	12	35		
Lymphoid tissue hyperplasia	10		5	3	5		
Lymphopenia	299	39	127	47	172	1	2
Macrocytosis	12	4	6	4	6		
Mast cell activation syndrome	90	46	90				1
Mastocytosis	14	2	5	3	9		1
Methaemoglobinaemia	1		1				
Microangiopathic haemolytic anaemia	8	2	7	1	1		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Blood and lymphatic system disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Microcytic anaemia	46	9	22	11	24		
Microcytosis	7	1	1		6		
Monoclonal B-cell lymphocytosis	3	2	3				
Monocytopenia	5		1		4		
Monocytosis	40	3	15	8	25		1
Myelocytosis	4		1	2	3		
Myelosuppression	17	5	17				
Necrotic lymphadenopathy	11	1	11				
Nephrogenic anaemia	3	2	3				
Neutropenia	258	86	258			1	3
Neutropenia neonatal	1		1				
Neutrophilia	89	10	45	14	44		
Normochromic anaemia	14	5	11	1	3		
Normochromic normocytic anaemia	15	2	6	3	9		
Normocytic anaemia	52	7	29	4	23		
Nucleated red cells	1				1		
Pancytopenia	154	43	154				
Paratracheal lymphadenopathy	13		4	3	9		
Pernicious anaemia	9	3	5	1	4		
Placental transfusion syndrome	2		2				
Plasma cell disorder	1				1		
Platelet anisocytosis	6		4		2		
Platelet destruction increased	1	1	1				
Platelet disorder	53	7	22	11	31		
Platelet dysfunction	3	1	1		2		
Platelet production decreased	1		1				
Poikilocytosis	6	2	2	1	4		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Blood and lymphatic system disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Polychromasia	7		2	2	5		
Polycythaemia	34	8	16	7	18		
Polycythaemia neonatorum	1		1				
Post-anaphylaxis mast cell anergy	1		1				
Pseudolymphoma	24	2	9	8	15		
Purpura non-thrombocytopenic	16	1	7	2	9		
Red blood cell abnormality	16		4	2	12		1
Reticulocytosis	4		2		2		
Reticuloendothelial dysfunction	1		1				
Retroperitoneal lymphadenopathy	3		2	1	1		
Sarcoidosis of lymph node	1	1	1				
Schistocytosis	1		1				
Secondary thrombocytosis	2	1	2				
Sickle cell anaemia with crisis	27	7	27				
Spherocytic anaemia	1				1		
Spleen atrophy	1		1				
Spleen congestion	2			2	2		
Spleen disorder	26	8	14	5	12		
Spleen ischaemia	3	2	3				
Splenic artery thrombosis	9	3	9				
Splenic cyst	6	1	1		5		
Splenic embolism	1		1				
Splenic granuloma	2	2	2				
Splenic haematoma	1	1	1				
Splenic haemorrhage	3	2	3				
Splenic induration	2	1	2				
Splenic infarction	82	23	82				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Blood and lymphatic system disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Splenic lesion	6	4	5		1		
Splenic necrosis	1		1				
Splenic thrombosis	7	3	7				
Splenic vein thrombosis	21	5	21				1
Splenitis	5		4		1		
Splenomegaly	206	43	113	44	93		
Spontaneous haematoma	214	19	64	47	150		2
Spontaneous haemorrhage	10	5	10			1	1
Spur cell anaemia	1			1	1		
Stress polycythaemia	1		1				
Thrombocytopenia	2002	485	2002			3	12
Thrombocytopenia neonatal	1	1	1				
Thrombocytopenic purpura	115	17	100	5	15	1	1
Thrombocytosis	109	20	57	26	52		
Thrombosis with thrombocytopenia syndrome	82	35	82				
Thrombotic microangiopathy	35	11	35				
Thrombotic thrombocytopenic purpura	110	30	110				1
Thymus disorder	8	1	3	2	5		
Thymus enlargement	5	2	2		3		
Warm autoimmune haemolytic anaemia	10	5	10				
White blood cell disorder	49	3	15	11	34	1	2
Total:	100970	5679	20806	32583	80164	51	169

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Abnormal precordial movement	1				1		
Accelerated idioventricular rhythm	1	1	1				
Accessory cardiac pathway	1	1	1				
Acquired left ventricle outflow tract obstruction	1		1				
Acute cardiac event	13	3	13				
Acute coronary syndrome	404	141	403	1	1	1	1
Acute left ventricular failure	11	4	11				
Acute myocardial infarction	1135	360	1135			2	2
Acute right ventricular failure	2	1	2				
Adams-Stokes syndrome	3		3				
Angina pectoris	4085	2048	4085			31	44
Angina unstable	72	21	72				1
Aortic valve calcification	6		3		3		
Aortic valve disease	10	3	7	2	3		
Aortic valve disease mixed	2	1	2				
Aortic valve incompetence	87	33	87				
Aortic valve sclerosis	8	2	5	1	3		
Aortic valve stenosis	26	6	22	1	4		
Aortic valve thickening	1		1				
Arrhythmia	9658	5568	9657	1	1	58	93
Arrhythmia neonatal	3	2	3			1	2
Arrhythmia supraventricular	83	25	62	9	21		
Arrhythmic storm	4	1	4				
Arteriosclerosis coronary artery	92	24	81		11		
Arteriospasm coronary	37	15	37				
Arteritis coronary	3	3	3				
Athletic heart syndrome	1			1	1		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Atrial enlargement	7	2	3	2	4		
Atrial fibrillation	3554	1142	3554			26	43
Atrial flutter	359	79	246	67	113	3	4
Atrial hypertrophy	1			1	1		
Atrial tachycardia	94	18	59	18	35	3	4
Atrial thrombosis	25	5	25				
Atrioventricular block	155	49	130	12	25		2
Atrioventricular block complete	129	35	129			2	3
Atrioventricular block first degree	48	6	28	11	20		
Atrioventricular block second degree	67	18	57	5	10		
Atrioventricular dissociation	2		2				
Autoimmune myocarditis	4		4				
Bezold-Jarisch reflex	1				1		
Bifascicular block	2		2				
Bradyarrhythmia	14	1	10	1	4		
Bradycardia	1295	301	1295			4	12
Bradycardia foetal	7	2	7				
Bradycardia neonatal						2	2
Bundle branch block	17	2	7	1	10		
Bundle branch block bilateral	3		1	1	2		
Bundle branch block left	145	49	96	27	49		1
Bundle branch block right	173	28	96	29	77	1	1
Cardiac amyloidosis	8	4	8				
Cardiac aneurysm	28	8	28				
Cardiac arrest	1516	391	1516			2	6
Cardiac arrest neonatal	1		1				
Cardiac asthma	13	1	13				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Cardiac contractility decreased	6	4	5	1	1		
Cardiac discomfort	1241	188	354	563	887	3	4
Cardiac disorder	1570	290	759	356	811	4	10
Cardiac dysfunction	91	28	91				
Cardiac failure	1909	573	1908	1	1	7	11
Cardiac failure acute	249	57	249				1
Cardiac failure chronic	96	37	96				
Cardiac failure congestive	249	70	249				3
Cardiac failure high output	1	1	1				
Cardiac fibrillation	188	78	188			2	3
Cardiac flutter	1196	349	1196			3	10
Cardiac granuloma	1	1	1				
Cardiac hypertrophy	64	17	47	8	17		
Cardiac perforation	3	2	3				
Cardiac perfusion defect	1		1				
Cardiac sarcoidosis	7	3	7				
Cardiac septal hypertrophy	5	1	2	1	3		
Cardiac steatosis	4	3	4				
Cardiac tamponade	108	37	108				
Cardiac valve discolouration	1	1	1				
Cardiac valve disease	56	12	33	13	23		
Cardiac valve sclerosis	3	1	3				
Cardiac ventricular disorder	8	1	3	3	5		
Cardiac ventricular thrombosis	37	9	37				
Cardiogenic shock	229	71	228	1	1		
Cardiomegaly	376	73	235	65	141	2	2
Cardiomyopathy	288	115	287		1	1	1

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Cardiomyopathy acute	2		2				
Cardiomyopathy alcoholic	1		1				
Cardiomyopathy neonatal	3	1	3				
Cardiopulmonary failure	80	19	80				
Cardiorenal syndrome	8	4	8				
Cardio-respiratory arrest	757	139	757				
Cardio-respiratory distress	9	3	9			1	1
Cardiovascular deconditioning	19	1	1	15	18		
Cardiovascular disorder	2404	204	618	1001	1786	3	6
Cardiovascular insufficiency	105	48	105				2
Cardiovascular symptom	35	9	24	5	11		
Carditis	178	112	178			1	2
Chordae tendinae rupture	4	3	4				
Chronic coronary syndrome	3	2	3				
Chronic left ventricular failure	1	1	1				
Conduction disorder	25	4	12	7	13		
Congestive cardiomyopathy	142	64	142				
Coronary artery aneurysm	4	2	4				
Coronary artery dilatation	10	3	6	3	4		
Coronary artery disease	261	81	204	29	57	5	5
Coronary artery dissection	25	10	25				
Coronary artery embolism	5	1	5				
Coronary artery insufficiency	3	1	3				
Coronary artery occlusion	106	31	106			1	3
Coronary artery perforation	1		1				
Coronary artery stenosis	108	25	108				
Coronary artery thrombosis	127	35	127				1

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Coronary ostial stenosis	4	2	4				
Cor pulmonale	19	4	19				
Cor pulmonale acute	36	15	36				
Cor pulmonale chronic	2	2	2				
Defect conduction intraventricular	14	2	8	3	6		
Degenerative aortic valve disease	2	1	2				
Degenerative mitral valve disease	1		1				
Diastolic dysfunction	43	7	22	8	21		
Dilatation atrial	16	4	10	2	6		
Dilatation ventricular	18	3	12		6		
Dressler's syndrome	8	1	5	2	3		
Early repolarisation syndrome	2	2	2				
Endocardial disease	1		1				
Endocardial fibrosis	2	1	2				
Endocarditis fibroplastica	1	1	1				
Endocarditis noninfective	1		1				
Endocarditis rheumatic	1	1	1				
Eosinophilic myocarditis	8	4	8				
Extrasystoles	1914	279	747	568	1167	5	8
Foetal arrhythmia	1	1	1				
Foetal cardiac arrest	17	2	17				
Foetal heart rate acceleration abnormality	1		1				
Foetal heart rate deceleration abnormality	1	1	1			1	1
Foetal heart rate disorder	1		1				
Gastrocardiac syndrome	4			1	4		
Giant cell myocarditis	2	2	2				
Haemorrhage coronary artery	3	3	3				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Heart alternation	15	3	5	6	10		
Heart valve calcification	5	1	4		1		
Heart valve incompetence	37	8	22	8	15		
Hepatojugular reflux	8		2	2	6		
Hyperdynamic left ventricle	2		2				
Hyperkinetic heart syndrome	2	2	2				
Hypersensitivity myocarditis	5	2	5				
Hypertensive cardiomyopathy	7	4	7				
Hypertensive heart disease	53	11	39	5	14		
Immune-mediated myocarditis	2	1	2				
Interventricular septum rupture	1		1				
Intracardiac mass	1		1				
Intracardiac thrombus	88	19	88				
Intrapericardial thrombosis	2	1	2				
Ischaemic cardiomyopathy	24	6	24				
Kounis syndrome	4	2	4				
Left atrial dilatation	20	2	9	4	11		
Left atrial enlargement	33	7	15	10	18		
Left ventricular dilatation	25	2	15	6	10		
Left ventricular dysfunction	162	51	122	16	40		
Left ventricular enlargement	9	3	7		2		
Left ventricular failure	87	32	87				2
Left ventricular hypertrophy	108	14	61	19	47		
Long QT syndrome	8	5	8				
Low cardiac output syndrome	8	4	8				
Lupus endocarditis	1	1	1				
Microvascular coronary artery disease	9	6	9				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Mitral valve calcification	7		4	1	3		
Mitral valve disease	21	4	12	4	9		
Mitral valve incompetence	235	101	235			6	6
Mitral valve prolapse	45	17	29	7	16		
Mitral valve sclerosis	4	3	4				
Mitral valve stenosis	6		6				
Mitral valve thickening	2		2				
Myocardial calcification	1				1		
Myocardial fibrosis	65	7	49	5	16		
Myocardial haemorrhage	3	2	3				
Myocardial hypoxia	10	5	10				
Myocardial infarction	2741	891	2741			29	38
Myocardial injury	116	46	116				
Myocardial ischaemia	239	71	239				1
Myocardial necrosis	31	9	31				1
Myocardial oedema	57	21	57				
Myocardial rupture	22	4	22				
Myocarditis	10363	4624	10363			15	26
Myocarditis post infection	2	1	2				
Myopericarditis	1792	692	1792			5	10
Nodal arrhythmia	4	1	3		1		
Nodal rhythm	11	1	8	1	3		
Palpitations	30435	4200	10639	8828	19796	30	74
Papillary muscle disorder	1		1				
Papillary muscle haemorrhage	1	1	1				
Papillary muscle rupture	1		1				
Paroxysmal arrhythmia	11	1	7	2	4		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Paroxysmal atrioventricular block	6		6				
Pericardial cyst	3			1	3		
Pericardial disease	46	13	27	8	19		
Pericardial effusion	1734	706	1734			10	14
Pericardial fibrosis	37	18	37				
Pericardial haemorrhage	42	13	42				
Pericardial mass	3	2	3				
Pericardial rub	29	4	14	9	15		
Pericarditis	9783	4114	9783			20	42
Pericarditis adhesive	1		1				
Pericarditis constrictive	16	5	16				
Pericarditis lupus	1	1	1				
Pericarditis rheumatic	1		1				
Pericarditis uraemic	1		1				
Pleuropericarditis	72	26	72				3
Pneumopericardium	2	1	2				
Positive cardiac inotropic effect	2			2	2		
Postinfarction angina	1		1				
Postural orthostatic tachycardia syndrome	306	72	178	78	128		1
Prinzmetal angina	36	17	36				1
Pulmonary valve disease	2		1		1		
Pulmonary valve incompetence	7	5	7				
Pulmonary valve stenosis	3	1	3				
Pulmonary valve thickening	1		1				
Pulseless electrical activity	40	12	40				
Rebound tachycardia	1				1		
Restrictive cardiomyopathy	2		2				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Rheumatic heart disease	5	4	5				
Rhythm idioventricular	1		1				
Right atrial dilatation	6		4	1	2		
Right atrial enlargement	10	1	3	3	7		
Right atrial hypertrophy	2	1	1	1	1		
Right ventricular diastolic collapse	1		1				
Right ventricular dilatation	27	8	17	2	10		
Right ventricular dysfunction	32	11	27	1	5		
Right ventricular enlargement	6	2	3	1	3		
Right ventricular failure	71	23	71				
Right ventricular hypertrophy	10	4	8	1	2		1
Silent myocardial infarction	6	4	6				
Sinoatrial block	8		8				
Sinus arrest	16	4	16				
Sinus arrhythmia	113	28	63	31	50		
Sinus bradycardia	148	31	88	21	60		1
Sinus node dysfunction	35	18	35				
Sinus tachycardia	849	152	438	162	411	1	2
Stress cardiomyopathy	121	36	106	7	15	1	3
Subendocardial haemorrhage	1		1				
Subendocardial ischaemia	2	1	2				
Supraventricular extrasystoles	296	55	151	50	145	3	3
Supraventricular tachyarrhythmia	8	2	5	1	3		
Supraventricular tachycardia	445	90	303	67	142		2
Systolic dysfunction	28	10	20	5	8		
Tachyarrhythmia	189	38	108	32	81	3	5
Tachycardia	25509	3002	7814	7876	17695	26	70

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Tachycardia foetal	11	2	11				1
Tachycardia induced cardiomyopathy	5	3	5				
Tachycardia paroxysmal	80	14	45	12	35		1
Torsade de pointes	14	4	14				
Toxic cardiomyopathy	4	3	4				
Tricuspid valve disease	5	2	5				
Tricuspid valve incompetence	117	44	117				
Tricuspid valve prolapse	1	1	1				
Trifascicular block	5	4	5				
Ventricle rupture	5	2	5				
Ventricular arrhythmia	68	24	68				
Ventricular asystole	1		1				
Ventricular dysfunction	38	8	28	3	10		
Ventricular dyskinesia	6	2	4		2		
Ventricular enlargement	10	5	7		3		
Ventricular extrasystoles	830	154	426	148	404	1	2
Ventricular failure	8	3	8				
Ventricular fibrillation	288	92	288			2	3
Ventricular flutter	4	2	4				
Ventricular hypertrophy	25	2	15	5	10		
Ventricular hypokinesia	127	47	127				
Ventricular pre-excitation	3	2	2	1	1		
Ventricular remodelling	2	1	2				
Ventricular tachyarrhythmia	5	2	5				
Ventricular tachycardia	284	98	284				
Wandering pacemaker	1		1				
Wolff-Parkinson-White syndrome	17	7	17				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Cardiac disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Total:	126993	33519	82303	20300	44690	327	608

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Aberrant aortic arch	2		2				
Abnormal palmar/plantar creases	1		1				
Accessory muscle	3			1	3		
Accessory spleen	4	1	2	1	2		
Acquired gene mutation	1		1				
Acral peeling skin syndrome	2	1	1		1		
Acrodermatitis enteropathica	1				1		
Adenomatous polyposis coli	2	1	1		1		
Adrenogenital syndrome	1		1				
Adrenoleukodystrophy	1			1	1		
Alpha-1 antitrypsin deficiency	2		1		1		
Amegakaryocytic thrombocytopenia	2		2				
Amyotrophic lateral sclerosis gene carrier	2		2				
Anal atresia	1	1	1				
Anencephaly	7		7				
Ankyloglossia congenital	3	1	3			1	1
Anodontia	1			1	1		
Anomalous pulmonary venous connection	2		2				
Anophthalmos	1		1				
Anotia	1		1				
Antithrombin III deficiency	6	1	5	1	1		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Aorta hypoplasia	1	1	1				
Aplasia	2	1	2				
Aplasia cutis congenita	1		1				
Arachnodactyly	1				1		
Argininosuccinate synthetase deficiency	2			1	2		
Arnold-Chiari malformation	7	1	7				
Arrhythmogenic right ventricular dysplasia	4		4				
Arterial tortuosity syndrome	1			1	1		
Arteriovenous malformation	8	4	8			2	2
ASAH1 related disorder	1		1				
Asplenia	1		1				
Asymmetric crying facies	1			1	1		
Ataxia telangiectasia	2		1	1	1		
Atrial septal defect	40	11	26	9	14	1	1
Atrioventricular septal defect	5	3	5				
Auditory neuropathy spectrum disorder	1	1	1				
Autoimmune lymphoproliferative syndrome	3	2	3				
Benign familial haematuria	1				1		
Benign familial pemphigus	9		1	2	8		
Bicuspid aortic valve	4	1	3	1	1		
Biliary hamartoma	1	1	1				
Birth mark	7		1	2	6		
Blindness congenital	1	1	1				
Block vertebra	5		2	2	3		
Brachydactyly	2	2	2				
Brachyolmia	1		1				
BRAF gene mutation	1	1	1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Brain malformation	2	2	2				
Branchial cyst	4	1	3	1	1		
Breast malformation	1		1				
Brugada syndrome	15	9	15				
CADASIL	1			1	1		
Cancer gene carrier	1				1		
Cardiac septal defect	2	2	2				
Camitine palmitoyltransferase deficiency	1		1				
Cataract congenital	2	1	2				
Caudal regression syndrome	1		1				
Cerebellar hypoplasia	1	1	1				
Cerebral arteriovenous malformation haemorrhagic	1	1	1				
Cerebral cavernous malformation	3	2	3				
Cerebral palsy	17	4	17				
Cerebrovascular arteriovenous malformation	4	3	4				
Chimerism	1				1		
Chordee	2			2	2		
Chronic granulomatous disease	1	1	1				
Cleft lip	2	2	2				
Cleft lip and palate	4	1	4				
Cleft palate	4	3	4				
Cleft uvula	1		1				
Clinodactyly	2		1		1		
Cloacal exstrophy	3	2	3				
Coarctation of the aorta	3		3				
Colour blindness	23	3	13	2	10		
Combined immunodeficiency	2	2	2				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Congenital absence of cranial vault	2		2				
Congenital amputation	1	1	1				
Congenital anomaly	45	16	44	1	1	2	3
Congenital aortic stenosis	1		1				
Congenital arterial malformation	1		1				
Congenital cardiovascular anomaly	1	1	1				
Congenital central nervous system anomaly	6	2	6				
Congenital cerebrovascular anomaly	1		1				
Congenital cystic kidney disease	2		2				1
Congenital cystic lung	3	2	3				
Congenital cytomegalovirus infection	1	1	1				
Congenital diaphragmatic hernia	1	1	1				
Congenital eye disorder	2	1	2				
Congenital foot malformation	1	1	1			1	1
Congenital genital malformation							1
Congenital great vessel anomaly	2		2				
Congenital haematological disorder	1	1	1				
Congenital hand malformation	1	1	1				
Congenital hearing disorder							1
Congenital heart valve disorder	2	1	2				
Congenital hydrocephalus	6	1	6				
Congenital hydronephrosis	3	2	3				
Congenital hypercoagulation	1		1				
Congenital inguinal hernia							1
Congenital jaw malformation	1	1	1				
Congenital LUMBAR syndrome	2	1	2				
Congenital methaemoglobinaemia	1		1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Congenital mitral valve incompetence	1	1	1				
Congenital musculoskeletal disorder	1	1	1				1
Congenital musculoskeletal disorder of limbs	7	4	7				1
Congenital musculoskeletal disorder of skull	2	1	2				
Congenital musculoskeletal disorder of spine	5	1	5				
Congenital myasthenic syndrome	1				1		
Congenital myopathy	1		1				
Congenital neurological degeneration	1	1	1				
Congenital nose malformation	1	1	1				
Congenital nystagmus	1			1	1		
Congenital optic nerve anomaly	1	1	1				
Congenital oral malformation	1	1	1				
Congenital skin dimples	1	1	1				
Congenital thyroid disorder							1
Congenital tricuspid valve atresia	1	1	1				
Comeal dystrophy	7	2	7				
Cortical dysplasia	2	1	2				
Craniosynostosis	2		2				
Cryopyrin associated periodic syndrome	2	1	2				
Cryptorchism	3	2	2			1	
CYP2C19 polymorphism	1					1	
Cystic fibrosis	1					1	
Cystic lymphangioma	7	2	7				
Cytogenetic abnormality	6	2	5	1	1		
Dacryostenosis congenital	2	1	1			1	
Dermoid cyst	7	1	7				
Developmental hip dysplasia	1		1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Diastematomyelia	1		1				
Diastrophic dysplasia	1		1				
Dolichocolon	1			1	1		
Double inlet left ventricle	1		1				
Double outlet right ventricle	2	2	2				
Ductus arteriosus premature closure	1		1				1
Dysmorphism	6	1	3		3		
Ectopic thyroid	1	1	1				
Ectrodactyly	1	1	1				
Ehlers-Danlos syndrome	20	3	9	7	11		
Elliptocytosis hereditary	1		1				
Encephalocele	3		3				
Epidermolysis bullosa	3	1	3				
Exomphalos	4	3	4				
Factor II mutation	4	1	1	1	3		
Factor IX deficiency	1	1	1				
Factor V deficiency	5	1	3	1	2		
Factor VII deficiency	1	1	1				
Factor VIII deficiency	9	3	8		1		
Factor V Leiden mutation	17	1	6	5	11		
Factor X deficiency	2		1	1	1		
Factor XI deficiency	1		1				
Factor XIII deficiency	1	1	1				
Falot's tetralogy	7	3	7				
Familial amyotrophic lateral sclerosis	2		2				
Familial hemiplegic migraine	2		2				
Familial mediterranean fever	7	1	3	2	4		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Familial periodic paralysis	5	1	3	1	2		
Familial tremor	2		1	1	1		
Fibrodysplasia ossificans progressiva	1	1	1				
Foetal chromosome abnormality	4	3	4				
Foetal cystic hygroma	6	1	6				
Foetal malformation	22	10	22				
Foramen magnum stenosis	1		1				
Friedreich's ataxia	1		1				
Gastrointestinal arteriovenous malformation						1	1
Gastrointestinal malformation	1		1				
Gastroschisis	4	1	4				
Gene mutation	10	1	3	6	7		
Gilbert's syndrome	21	2	8	5	13		
Glucose-6-phosphate dehydrogenase deficiency	4		1		3		
Glycogen storage disease type V	4	2	4				
Glycogen storage disorder	1				1		
Grey matter heterotopia	1	1	1				
Haemophilia	12	3	8	2	4		
Haemorrhagic arteriovenous malformation	1		1				
Hamartoma	2	1	1	1	1		
Heart disease congenital	40	15	40				2
Hereditary angioedema	9	6	8		1		1
Hereditary angioedema with normal C1 esterase inhibitor	1		1				
Hereditary ataxia	3	2	3				
Hereditary fructose intolerance	2			2	2		
Hereditary haemorrhagic telangiectasia	4			2	4		
Hereditary motor and sensory neuropathy	5	2	3		2		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Hereditary neuropathy with liability to pressure palsies	3		3				
Hereditary optic atrophy	1		1				
Hereditary spherocytosis	2		1	1	1		
Heterotaxia	2		2				
Homocystinaemia	1			1	1		
Huntington's disease	10	2	7	1	3		
Hydranencephaly	1		1				
Hydrocele	13	2	4	4	9		
Hyperreflexia	3		1		2		
Hyperglycinaemia	3		2		1		
Hyper IgD syndrome	1				1		
Hypertrophic cardiomyopathy	43	17	43				
Hypochondroplasia	1	1	1				
Hypophosphatasia	1	1	1				
Hypoplastic left heart syndrome	4	1	4				
Hypospadias					1	2	
Ichthyosis	1	1	1				
Imperforate oesophagus	1		1				
Inborn error of metabolism	1			1	1		
Intestinal atresia	4	3	4			1	1
Intracranial lipoma	1		1				
Isodicentric chromosome 15 syndrome	1	1	1				
Janus kinase 2 mutation	2	1	2				
Keams-Sayre syndrome	1				1		
Keratosis follicular	4	2	3		1		
Kidney duplex	2		1	1	1		
Kidney malformation	5	3	5				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Klippel-Trenaunay syndrome	1		1				
Laryngomalacia	2	1	2				
Left-to-right cardiac shunt	1		1				
Limb hypoplasia congenital	1		1				
Limb reduction defect	2		2				
Long QT syndrome congenital	1		1				
Low set ears	1		1				
Macrocephaly	1		1			1	1
Macroglossia	26	5	26				1
Malformation venous	1		1				
MELAS syndrome	1		1				
Melkersson-Rosenthal syndrome	4	2	4				
Meningocele	1		1				
Meningomyelocele	1	1	1				
Metabolic myopathy	1			1	1		
Methylenetetrahydrofolate reductase gene mutation	6		1	2	5		
Microcephaly	1		1				
Micrognathia	1		1				
Micropenis	4	1	4				
Microphthalmos	1	1	1				
Mitochondrial encephalomyopathy	1		1				
Mitochondrial myopathy	2			2	2		
Mitral valve atresia	2		2				
Moebius II syndrome	1		1				
Multiple congenital abnormalities	3	2	3				
Muscular dystrophy	9	1	2	4	7		
Myocardial bridging	3	1	3				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Myoclonic dystonia	4	1	4				
Myoclonic epilepsy and ragged-red fibres	1			1			
Myotonic dystrophy	2		1	1	1		
Naevus flammeus	3		1	2	2		
Nail aplasia	1		1				
Neonatal alloimmune thrombocytopenia	1		1				
Neonatal haemochromatosis	1		1				
Neurofibromatosis	5	1	3	1	2		
Non-compaction cardiomyopathy	2	2	2				
Odontogenic cyst	1			1	1		
Osteogenesis imperfecta	2	1	2				
Pachyonychia congenita	1	1	1				
Pancreas divisum	1				1		
Parkes-Weber syndrome	1				1		
Paroxysmal extreme pain disorder	4		2	2	2		
Patent ductus arteriosus						2	2
Pectus excavatum	2	1	1	1	1		
Penoscrotal fusion	1	1	1				
Persistent foetal circulation	1	1	1				
Pfeiffer syndrome	1				1		
Phelan-McDermid syndrome	1		1				
Phimosis	6	1	3	3	3		
Platybasia	1		1				
Polycystic liver disease	1	1	1				
Polydactyly	2	1	1		1		
Polymicrogyria	1		1				
Porencephaly	1		1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Porokeratosis	1		1				
Porphyria	4	1	4				
Porphyria acute	5	2	5				1
Porphyria non-acute	3	2	3				
Potter's syndrome	1	1	1				
Preauricular cyst	2		1		1		
Primary ciliary dyskinesia	1			1	1		
Protein C deficiency	3		1	2	2		
Protein S deficiency	5	3	5				
PTEN gene mutation	1		1				
Pulmonary artery stenosis congenital	1	1	1				
Pulmonary hypoplasia	2		2				
Pulmonary malformation	2	1	2			1	1
Pulmonary sequestration	2	2	2				
Pulmonary valve stenosis congenital	2	2	2			1	1
Rathke's cleft cyst	2	1	1	1	1		
Renal aplasia	2	2	2				
Renal dysplasia	1		1			1	2
Renal fusion anomaly	1	1	1				
Retinitis pigmentosa	6	1	6				
Right-to-left cardiac shunt	1		1				
Rippling muscle disease	1				1		
RUNX1 gene mutation	1	1	1				
Schizencephaly	1		1				
Scimitar syndrome	1		1				
Sensory neuropathy hereditary	2	1	2				
Short-chain acyl-coenzyme A dehydrogenase deficiency	1		1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Sickle cell anaemia	2		1	1	1		
Sickle cell disease	2	1	1		1		
Single atrium	1		1				
Single umbilical artery	5	1	4	1	1		
Skeletal dysplasia	1	1	1				
Skin malformation	2		2				
Spina bifida	6	2	6				
Spinal muscular atrophy	2	2	2				
Spinal vessel congenital anomaly	1		1				
Stargardt's disease	1			1	1		
Sturge-Weber syndrome	1		1				
Supernumerary nipple	3		1		2		
Syndactyly	4	1	4				
Syringomyelia	9	4	9				
Talipes	2		2				
Telangiectasia congenital	1		1				
Thalassaemia	1	1	1				
Thalassaemia minor	7	1	2	2	5		
Thyroglossal cyst	4		3	1	1		
Tourette's disorder	21	8	21				
Tracheo-oesophageal fistula	1		1				
Transposition of the great vessels	4	1	4				
Trisomy 13	2	1	2				
Trisomy 16	1	1	1				
Trisomy 18	4	1	4				
Trisomy 21	11	3	11				
Trisomy 8	1	1	1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Congenital, familial and genetic disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Turner's syndrome	3	1	1	1	2		
Type IIa hyperlipidaemia	2		1		1		
Type V hyperlipidaemia	7		3		4		
Univentricular heart	1	1	1				
Urachal abnormality	1	1	1				
Urethral atresia	1		1				
Urethral valves	3	2	3				
Urinary tract malformation							1
VACTERL syndrome	1	1	1				
Vascular endothelial growth factor overexpression	1			1	1		
Vascular malformation	6	1	2	3	4		
Venous angioma of brain	4	1	4				
Ventricular hypoplasia	2		2				
Ventricular septal defect	14	6	14			1	1
Vertebral artery hypoplasia	1		1				
Vestibulocerebellar syndrome	3		2		1		
VEXAS syndrome	1	1	1				
Von Willebrand's disease	5	2	5				
Williams syndrome	1		1				
Wolf-Hirschhorn syndrome	1		1				
X-linked lymphoproliferative syndrome	1	1	1				
XXX syndrome	1		1				
Total:	1143	345	904	116	239	17	34

System Organ Class

Ear and labyrinth disorders

Spontaneous		Non Interventional Study
Serious	Nonserious	Serious

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Ear and labyrinth disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Acute vestibular syndrome	80	20	51	8	29		
Auditory disorder	238	20	62	79	176		
Aural polyp	1	1	1				
Auricular swelling	42	1	11	10	31		
Autoimmune inner ear disease	2	1	2				
Autophony	4		2	1	2		
Cerumen impaction	10		1	3	9		
Chondrodermatitis nodularis chronica heliis	1		1				
Conductive deafness	9	2	9				
Deafness	1448	484	1448			5	11
Deafness bilateral	99	27	99				
Deafness neurosensory	220	65	220				
Deafness permanent	2	2	2			1	1
Deafness transitory	68	21	68				
Deafness unilateral	719	222	719			2	4
Diplacusis	10		4	2	6		
Dysacusis	25	2	9	5	16		
Ear canal erythema	11	1	3		8		
Ear canal stenosis	3	1	1	2	2		
Ear congestion	219	11	52	54	167		
Ear deformity acquired	3				3		
Ear discomfort	2217	135	508	508	1709	1	1
Ear disorder	286	24	88	53	198	1	1
Ear haemorrhage	90	11	29	22	61		
Ear inflammation	61	15	20	32	41		
Ear odour	1				1		
Ear pain	5188	336	1454	1084	3734	3	11

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Ear and labyrinth disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Ear pruritus	242	7	51	32	191		1
Ear swelling	388	20	103	81	285		1
Endolymphatic hydrops	19	8	19				
Eustachian tube disorder	13	1	9	2	4		
Eustachian tube dysfunction	35	7	12	6	23		
Eustachian tube obstruction	32	5	9	6	23		
Eustachian tube patulous	2				2		
Excessive cerumen production	42	3	13	7	29		1
External ear disorder	5		4	1	1		
External ear inflammation	16		6	3	10		
External ear pain	42	2	13	6	29		
Haematotympanum	1		1				
Hyperacusis	847	105	301	210	546		1
Hypoacusis	2096	255	772	490	1324	2	4
Inner ear disorder	85	10	37	15	48		
Inner ear infarction	8	3	8				
Inner ear inflammation	40	3	19	7	21		
Mastoid disorder	5		1	1	4		
Mastoid effusion	1				1		
Meniere's disease	193	53	193				
Middle ear disorder	6	1	3	2	3		
Middle ear effusion	36	3	9	11	27		1
Middle ear inflammation	36	5	7	20	29		
Misophonia	12		3	2	9		
Mixed deafness	4	2	4				
Motion sickness	203	18	86	25	117		1
Neonatal deafness	1	1	1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Ear and labyrinth disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Neurosensory hypoacusis	38	6	27	4	11	1	2
Noninfective myringitis	5			3	5		
Otolithiasis	22		8	8	14		
Otorrhoea	56	7	16	13	40		
Otosclerosis	8	3	6	1	2		
Ototoxicity	5	4	5				
Paraesthesia ear	41	1	12	8	29		
Phobic postural vertigo	17	4	5	6	12		
Presbycusis	9	2	3	5	6		
Red ear syndrome	10		1	1	9		
Sudden hearing loss	882	363	882			7	8
Superior semicircular canal dehiscence	1		1				
Tinnitus	15833	1522	4286	4532	11547	11	30
Tympanic membrane disorder	28	4	8	4	20		
Tympanic membrane hyperaemia	5	2	3	1	2		
Tympanic membrane perforation	27	5	13	8	14		
Vertigo	13698	964	3909	2713	9789	18	56
Vertigo labyrinthine	23	7	17	1	6		
Vertigo positional	650	79	259	152	391	3	5
Vestibular ataxia	6	2	6				
Vestibular disorder	203	36	103	34	100	1	1
Vestibular ischaemia	3	1	3				
Vestibular paroxysmia	1		1				
Total:	47038	4926	16122	10284	30916	56	141

System Organ Class

Endocrine disorders

Spontaneous	Non Interventional Study
-------------	--------------------------

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Endocrine disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Addison's disease	10	7	10				
Adrenal atrophy	1			1	1		
Adrenal cortex necrosis	2		2				
Adrenal disorder	26	5	11	5	15		
Adrenal haemorrhage	10	2	10				
Adrenal insufficiency	55	21	55				
Adrenal mass	9	1	1	4	8		
Adrenal thrombosis	1		1				
Adrenocortical insufficiency acute	56	16	56				
Adrenocorticotrophic hormone deficiency	1		1				
Adrenomegaly	7		1	1	6		
Androgen deficiency	2				2		
Anovulatory cycle	294	29	69	127	225		
Autoimmune hypothyroidism	8	6	8				
Autoimmune thyroid disorder	14	8	14				
Autoimmune thyroiditis	336	194	336			4	4
Basedow's disease	418	221	418			1	2
Carcinoid crisis	1	1	1				
Central hypothyroidism	2	1	2				
Cushingoid	8	1	3	1	5		
Cushing's syndrome	4	1	4				
Delayed menarche	7	2	2		5		
Diabetes insipidus	29	20	29				
Empty sella syndrome	2	2	2				
Endocrine disorder	53	4	21	17	32	1	1
Euthyroid sick syndrome	2				2		
Fertility increased	3			3	3		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Endocrine disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Glucocorticoid deficiency	3	2	3				
Goitre	232	29	88	64	144		
Growth hormone deficiency	2	1	2				
Haemorrhagic thyroid cyst	2			1	2		
Hyperadrenocorticism	1	1	1				
Hyperaldosteronism	2		2				
Hypercalcaemia of malignancy	1				1		
Hyperoestrogenism	1				1		
Hyperparathyroidism	8		3	1	5		
Hyperparathyroidism primary	5	1	4		1		
Hyperparathyroidism secondary	2	1	2				
Hyperplasia adrenal	3	3	3				
Hyperprolactinaemia	5	4	5				
Hyperthyroidism	546	245	546			3	7
Hypoparathyroidism	2		1		1		
Hypophysitis	5		2	2	3		
Hypopituitarism	14	1	6	6	8		
Hypothalamo-pituitary disorder	13	9	13				
Hypothyroidic goitre	2			1	2		
Hypothyroidism	433	220	433			2	4
Immune-mediated hypothyroidism	1		1				
Immune-mediated thyroiditis	3	2	3				
Inappropriate antidiuretic hormone secretion	23	9	23				
Luteal phase deficiency	6		1	4	5		
Lymphocytic hypophysitis	1			1	1		
Myxoedema	5	3	5				
Oestrogen deficiency	4		1	3	3		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Endocrine disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Ovarian dysfunction	1				1		
Ovulation delayed	196	4	25	81	171		
Parathyroid disorder	3	1	1	1	2		
Pituitary apoplexy	6	4	6				
Pituitary-dependent Cushing's syndrome	2	1	2				
Pituitary enlargement	2	1	1	1	1		
Pituitary haemorrhage	3		3				
Pituitary infarction	1		1				
Polyglandular autoimmune syndrome type I	1		1				
Polyglandular autoimmune syndrome type II	1		1				
Precocious puberty	2			2	2		
Premature menarche	27		4	3	23		
Primary hyperaldosteronism	3		1	1	2		
Primary hyperthyroidism	6	4	6				
Secondary adrenocortical insufficiency	4		3	1	1		
Secondary hyperthyroidism	2	2	2				
Secondary hypogonadism	1		1				
Silent thyroiditis	9	3	7	1	2		
Thyroid calcification	1				1		
Thyroid cyst	28	4	10	8	18		
Thyroid dermatopathy	1		1				
Thyroid disorder	299	40	82	120	217	3	3
Thyroid haemorrhage	1		1				
Thyroiditis	225	44	102	52	123	2	2
Thyroiditis acute	53	15	53			1	2
Thyroiditis chronic	3	1	1		2		
Thyroiditis subacute	286	65	143	59	143		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Endocrine disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Thyroid mass	123	19	45	41	78		
Thyroid pain	92	7	21	35	71		
Thyroid stimulating hormone deficiency	3			1	3		
Thyrotoxic crisis	34	15	34				
Thyrotoxic periodic paralysis	1			1	1		
Toxic goitre	3	2	3				
Toxic nodular goitre	6	5	6				
Total:	4115	1310	2772	650	1343	17	25

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Abnormal sensation in eye	219	6	52	38	167		
Accommodation disorder	65	6	15	19	50		
Acute macular neuroretinopathy	18	9	18				
Age-related macular degeneration	13	6	13				
Altered visual depth perception	22		5	4	17		
Amaurosis	29	5	29				
Amaurosis fugax	101	42	101			1	1
Amblyopia	11		4	2	7		
Anaesthesia eye	1			1	1		
Angle closure glaucoma	17	6	17				
Anisocoria	90	19	54	17	36		
Anisometropia	1				1		
Anterior chamber cell	1		1				
Anterior chamber flare	2		1	1	1		

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Anterior chamber inflammation	3	2	3				
Anterior chamber opacity	1			1			
Anterior segment ischaemia	1	1	1				
Arteriosclerotic retinopathy	3	1	3				
Asthenopia	676	43	219	99	457		3
Astigmatism	32	5	8	11	24		
Atopic keratoconjunctivitis	1		1				
Autoimmune eye disorder	3	1	2	1	1		
Autoimmune uveitis	6	4	6				
Bell's phenomenon	5	1	3	1	2		
Binocular eye movement disorder	15		8	5	7		
Birdshot chorioretinopathy	2		2				
Blepharal pigmentation	1			1	1		
Blepharitis	195	13	41	57	154		
Blepharitis allergic	2	1	1		1		
Blepharochalasis	3			2	3		
Blepharophimosis	1				1		
Blepharospasm	784	46	158	230	626		1
Blindness	970	318	970			2	4
Blindness cortical	5		5				
Blindness transient	274	91	274			1	1
Blindness unilateral	328	103	328			1	1
Cataract	150	72	150			12	19
Cataract cortical	1		1				
Cataract nuclear	4	1	4				
Cataract subcapsular	3	1	1	1	2		
Central serous chorioretinopathy	46	25	46				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Central vision loss	13	4	13				
Chalazion	55	2	11	14	44		
Charles Bonnet syndrome	3		2		1		
Chloropsia	3	1	3				
Chorioretinal disorder	4		2	1	2		
Chorioretinal folds	1	1	1				
Chorioretinopathy	30	14	29		1		
Choroidal detachment	2	2	2				
Choroidal effusion	4		4				
Choroidal haemorrhage	1	1	1				
Choroidal infarction	1	1	1				
Choroidal neovascularisation	10	4	10				
Choroidal rupture	2		2				
Choroiditis	7	2	7				
Chromatopsia	29	4	9	7	20		
Ciliary muscle spasm	3			1	3		
Cogan's syndrome	2	2	2				
Colour blindness acquired	2	2	2				
Computer vision syndrome	2				2		
Conjunctival adhesion	2		2				
Conjunctival cyst	3			1	3		
Conjunctival defect	1				1		
Conjunctival discolouration	3			1	3		
Conjunctival disorder	12	1	3	3	9		
Conjunctival haemorrhage	437	104	437				5
Conjunctival hyperaemia	222	12	89	27	133		1
Conjunctival irritation	28	1	6	4	22		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Conjunctival oedema	35	2	14	6	21		
Conjunctival pallor	6		2	2	4		
Conjunctival suffusion	1	1	1				
Conjunctival telangiectasia	1				1		
Conjunctival ulcer	2	2	2				
Conjunctival vascular disorder	1		1				
Conjunctivitis allergic	60	5	12	17	48		
Contact lens intolerance	1			1	1		
Corneal bleeding	11	5	11				
Corneal decompensation	1	1	1				
Corneal defect	1		1				
Corneal degeneration	2		2				
Corneal deposits	2			1	2		
Corneal disorder	24	8	16	3	8		
Corneal endothelial cell loss	1	1	1				
Corneal endotheliitis	3	2	3				
Corneal epithelium defect	2	2	2				
Corneal erosion	11	1	5	3	6		
Corneal exfoliation	1		1				
Corneal infiltrates	3		2		1		
Corneal irritation	1			1	1		
Corneal lesion	5		1	3	4		
Corneal oedema	11	2	5	3	6		
Corneal opacity	13	2	13			1	1
Corneal perforation	1	1	1				
Corneal pigmentation	1			1	1		
Corneal scar	1		1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Corneal thinning	1		1				
Cyanopsia	12	2	6	2	6		
Cyclitis	1	1	1				
Cystoid macular oedema	19	7	19				
Dacryoadenitis acquired	7	3	4	1	3		
Dacryostenosis acquired	6		1	1	5		
Dark circles under eyes	54	5	12	15	42		
Delayed dark adaptation	1			1	1		
Dellen	1				1		
Dermatochalasis	12	2	6	4	6		
Detached Descemet's membrane	1		1				
Detachment of retinal pigment epithelium	2		2				
Diabetic eye disease	1				1		
Diabetic retinal oedema	1				1		
Diabetic retinopathy	2		2				
Diplopia	1722	268	879	278	843	3	4
Disorder of orbit	3	1	1	2	2		
Dry age-related macular degeneration	6	2	6				1
Dry eye	958	67	245	234	713	3	6
Dyschromatopsia	57	10	32	7	25		
Dysmetropsia	2			1	2		
Ectropion	4		3	1	1		
Eczema eyelids	53	3	8	18	45		
Endocrine ophthalmopathy	35	21	35			1	2
Enophthalmos	1	1	1				
Epiretinal membrane	7	4	6	1	1		
Episcleritis	68	4	21	15	47		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Erythema of eyelid	229	10	74	36	155	1	2
Erythroptosis	8		2	2	6		
Excessive eye blinking	29	1	9	7	20		
Excessive ocular convergence	1		1				
Exfoliation syndrome	1				1		
Exophthalmos	50	20	50				
Exposure keratitis	2			1	2		
Extraocular muscle disorder	20		12	1	8		
Exudative retinopathy	1	1	1				
Eye allergy	76	7	23	16	53		
Eye colour change	24	2	10	3	14		
Eye discharge	218	12	62	39	156		
Eye disorder	855	69	283	169	572	2	7
Eye haematoma	65	17	65				
Eye haemorrhage	566	175	565	1	1	3	8
Eye infarction	50	20	50			1	1
Eye inflammation	571	47	125	171	446	1	3
Eye irritation	1531	71	262	263	1269		3
Eyelash changes	2		1		1		
Eyelash discolouration	6		1	2	5		
Eyelid bleeding	11	2	5	3	6		
Eyelid cyst	16	1	8	3	8		
Eyelid disorder	111	8	44	18	67		1
Eyelid exfoliation	10		2	1	8		
Eyelid function disorder	97	7	47	13	50		
Eyelid haematoma	50	5	13	6	37		
Eyelid irritation	124	7	25	28	99		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Eyelid margin crusting	19	1	3	2	16		
Eyelid myoclonus	10	3	10				
Eyelid myokymia	23	3	8	6	15		
Eyelid oedema	1004	43	275	187	729	1	4
Eyelid pain	166	6	35	31	131		1
Eyelid ptosis	485	76	246	66	239		2
Eyelid rash	76	8	25	13	51		
Eyelid retraction	6		3		3		
Eyelid sensory disorder	35	2	8	6	27		
Eyelid skin dryness	18		3	5	15		
Eyelids pruritus	149	2	38	17	111		
Eyelid thickening	14		3	4	11		
Eyelid vascular disorder	4	1	1		3		
Eye movement disorder	353	49	201	57	152		1
Eye oedema	228	13	68	40	160		2
Eye opacity	2			1	2		
Eye pain	6217	447	1821	1288	4396	2	17
Eye paraesthesia	57	2	21	7	36		
Eye pruritus	1680	60	392	281	1288	4	7
Eye swelling	2929	185	866	622	2063	2	6
Eye symptom	28	5	11	4	17		
Eye ulcer	15	3	8	3	7		
Floppy eyelid syndrome	2		2				
Foreign body sensation in eyes	124	13	35	31	89		
Fuchs' syndrome	4	1	4				1
Gaze palsy	99	27	98		1		
Giant papillary conjunctivitis	3		1		2		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Glare	33	3	9		24		
Glaucoma	107	38	107			6	10
Glaucomatocyclitic crises	2		2				
Growth of eyelashes	1		1				
Halo vision	35	4	16	3	19		
Heteronymous diplopia	1		1				
Heterophoria	10	2	5	2	5		
Holmes-Adie pupil	8	1	2	5	6		
Homonymous diplopia	4	2	4				
Hyperaesthesia eye	6	1	2		4		
Hypermetropia	27	5	10	8	17		
Hypoaesthesia eye	128	15	49	18	79		
Hypotony of eye							1
Idiopathic orbital inflammation	5	2	5				
Inflammation of lacrimal passage	2	1	1	1	1		
Iridocele	1	1	1				
Iridocyclitis	161	62	161				1
Iris adhesions	4	3	4				
Iris cyst	1			1	1		
Iris discolouration	2			1	2		
Iris disorder	6		1	4	5		
Iris neovascularisation	1	1	1				
Iritis	119	22	52	34	67	2	2
Keratic precipitates	5	1	5				
Keratitis	86	13	37	18	49		
Keratitis interstitial	1	1	1				
Keratoconus	5	2	5			1	1

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Lacrimal disorder	12	1	2	4	10		
Lacrimal gland enlargement	5	1	2	1	3		
Lacrimal structural disorder	1		1				
Lacrimation decreased	16	4	10	4	6		
Lacrimation disorder	13		2	2	11		
Lacrimation increased	1555	85	341	326	1214	1	7
Lagophthalmos	43	7	26	4	17	1	1
Lens dislocation	1		1				
Lens disorder	4			3	4		
Lenticular opacities	4	1	4				
Lid margin discharge	1				1		
Lid sulcus deepened	12	5	7	2	5		
Limbal swelling	4		3		1		
Loss of visual contrast sensitivity	2		1		1		
Macular cherry-red spots	2	2	2				
Macular degeneration	54	14	54			2	3
Macular detachment	4	3	4				
Macular fibrosis	2		2				
Macular hole	13	5	13				2
Macular ischaemia	3	2	3				
Macular oedema	112	42	112				1
Macular opacity	1	1	1				
Macular rupture	2		2				
Macular scar	1				1		
Maculopathy	35	5	25	4	10		
Meibomian gland discharge	1				1		
Meibomian gland dysfunction	8		2	4	6		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Meibomianitis	2	1	1	1	1		
Metamorphopsia	90	12	51	12	39		
Miosis	57	5	22	9	35		
Mydriasis	246	20	100	31	146		
Myopia	61	10	21	19	40		
Myopic chorioretinal degeneration	1		1				
Necrotising retinitis	11	6	11				
Necrotising scleritis	1		1				
Neovascular age-related macular degeneration	24	12	24				
Neurological eyelid disorder	5		1	3	4		
Night blindness	16	1	2	6	14		
Noninfective chorioretinitis	2	1	2				
Noninfective conjunctivitis	2				2		
Ocular discomfort	1349	85	257	418	1092		
Ocular dysmetria	2		1		1		
Ocular hyperaemia	2176	104	522	416	1654	4	9
Ocular hypertension	74	39	74			1	1
Ocular ischaemic syndrome	7	3	7				
Ocular myasthenia	31	13	31				
Ocular rosacea	8	2	2	2	6		
Ocular sarcoidosis	2		2				
Ocular vascular disorder	106	23	85	3	21		
Ocular vasculitis	9	6	9				
Oculogyric crisis	17	6	17				1
Open angle glaucoma	4	1	4				
Ophthalmic artery aneurysm	1		1				
Ophthalmic artery occlusion	5	5	5				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Ophthalmic artery thrombosis	13	4	13				
Ophthalmic vein thrombosis	151	73	151			1	1
Ophthalmoplegia	125	43	125			1	1
Opsoclonus myoclonus	3	2	3				
Optic atrophy	13	6	13				
Optic disc disorder	7		3	3	4		
Optic disc haemorrhage	7	2	7				
Optic discs blurred	1		1				
Optic ischaemic neuropathy	150	47	150				
Optic nerve compression	1	1	1				
Optic nerve disorder	45	9	27	8	18		1
Optic nerve infarction	11	3	11				
Optic nerve sheath haemorrhage	1		1				
Optic neuropathy	34	12	34				
Orbital cyst	4	1	3		1		
Orbital haematoma	5		5				
Orbital haemorrhage	2		2				
Orbital myositis	5	3	5				
Orbital oedema	12	4	11	1	1		
Orbital swelling	12	1	2	3	10		
Oscillopsia	12	2	5	3	7		
Panophthalmitis	3	1	3				
Papilloedema	152	68	152				
Papillophlebitis	3	1	3				
Paralytic lagophthalmos	6	1	6				
Parophthalmia	6		2	4	4		
Periorbital discomfort	17		3	3	14		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Periorbital disorder	6				6		
Periorbital fat herniation	1			1	1		
Periorbital inflammation	3		1	1	2		
Periorbital irritation	2				2		
Periorbital oedema	301	36	124	66	177		4
Periorbital pain	56	9	24	11	32		1
Periorbital swelling	890	40	215	218	675		
Photokeratitis	1	1	1				
Photophobia	2800	194	949	547	1851	3	5
Photopsia	814	68	302	160	512		1
Pigmentary glaucoma	2		2				
Pinguecula	2	1	1		1		
Posterior capsule opacification	3	2	3				
Presbyopia	12	1	3	2	9		
Pseudomyopia	3		1	1	2		
Pseudopapilloedema	1			1	1		
Pterygium	4		1	1	3		
Punctate keratitis	12	4	12				
Pupil fixed	23	5	23				
Pupillary deformity	2		2				
Pupillary disorder	19	2	7	3	12		
Pupillary reflex impaired	34	4	21	5	13		
Recession of chamber angle of eye	1		1				
Refraction disorder	5	1	2	2	3		
Retinal aneurysm	7	3	7				
Retinal artery embolism	15	6	15				
Retinal artery occlusion	207	52	207				2

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Retinal artery stenosis	2	2	2				
Retinal artery thrombosis	32	5	29	1	3		
Retinal cyst	2	1	2				
Retinal degeneration	19	6	19				
Retinal depigmentation	1		1				
Retinal detachment	204	82	204			3	4
Retinal disorder	29	3	13	5	16		
Retinal drusen	3	1	3				
Retinal dystrophy	2		2				
Retinal exudates	16	7	15	1	1		
Retinal fibrosis	1		1				
Retinal fovea disorder	2		2				
Retinal haemorrhage	142	55	142			1	2
Retinal infarction	11	4	11				
Retinal ischaemia	18	8	18			1	1
Retinal microangiopathy	2	2	2				
Retinal neovascularisation	3	2	3				
Retinal oedema	26	11	26				
Retinal pigmentation	2		1	1	1		
Retinal pigment epitheliopathy	7	1	7				
Retinal scar	2		1		1		
Retinal tear	86	38	86				2
Retinal toxicity	3		3				
Retinal vascular disorder	17	9	17				
Retinal vascular occlusion	38	19	38				
Retinal vascular thrombosis	91	21	80	5	11		1
Retinal vasculitis	17	10	17				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Retinal vein occlusion	416	123	416			4	4
Retinal vein thrombosis	155	49	141	5	14		1
Retinal vein varices	1				1		
Retinal vessel avulsion	1		1				
Retinal white dots syndrome	12	10	12				
Retinopathy	21	5	21				
Retinopathy hypertensive	6	2	6				
Retinoschisis	3	1	3				
Rhegmatogenous retinal detachment	2	1	2				
Saccadic eye movement	5	1	1	3	4		
Scintillating scotoma	58	5	20	7	38		
Scleral cyst	4		1	1	3		
Scleral discolouration	14		4	2	10		
Scleral disorder	6		3	1	3		
Scleral haemorrhage	10		6	1	4		
Scleral hyperaemia	5	1	1	1	4		
Scleral oedema	3	3	3				
Scleral pigmentation	1			1	1		
Scleritis	51	13	51				
Serous retinal detachment	7	6	7				
Spontaneous hyphaema	2		2				
Staphyloma	1		1				
Strabismus	95	12	48	12	47		
Subretinal fluid	4	1	4				
Subretinal haematoma	2	1	2				
Sudden visual loss	36	11	36				
Swelling of eyelid	1307	47	278	297	1029		3

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Swollen tear duct	5		1		4		
Symblepharon	1		1				
Sympathetic ophthalmia	1		1				
Tear discolouration	1			1	1		
Tolosa-Hunt syndrome	7	3	7				
Tractional retinal detachment	1	1	1				
Ulcerative keratitis	23	11	23				
Uveitis	346	137	346			5	9
Venous stasis retinopathy	4		4				
Vision blurred	8600	894	3015	1659	5585	7	24
Visual acuity reduced	602	75	276	158	326		1
Visual acuity reduced transiently	18		2	7	16		
Visual brightness	29	1	12		17		
Visual field defect	632	104	331	115	301	1	3
Visual impairment	7540	846	2556	2238	4984	7	22
Visual snow syndrome	27	3	17	6	10		
Vitreoretinal traction syndrome	3	1	3				
Vitreous adhesions	5	4	4	1	1		
Vitreous degeneration	5	2	5				
Vitreous detachment	229	81	229			1	2
Vitreous disorder	24	2	11	6	13		
Vitreous floaters	527	54	197	97	330	1	1
Vitreous haematoma	1		1				
Vitreous haemorrhage	94	41	94				
Vitreous haze	10	6	10				
Vitreous opacities	30	6	9	18	21		
Vitreous prolapse	1	1	1				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Eye disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Vitritis	5	2	5				
Vogt-Koyanagi-Harada disease	15	7	15				
Xanthopsia	7		3	2	4		
Xerophthalmia	11	4	11				
Total:	61518	7145	24527	11690	36991	96	252

System Organ Class

Gastrointestinal disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Abdominal adhesions	7	2	6		1		
Abdominal discomfort	5454	235	1069	1212	4385	7	20
Abdominal distension	2791	194	805	600	1986	7	15
Abdominal fat apron	1				1		
Abdominal hernia	11	2	7		4		1
Abdominal incarcerated hernia	2		2				
Abdominal mass	27	2	9	3	18		
Abdominal migraine	4		1	1	3		
Abdominal pain	19593	1186	4277	4506	15316	19	57
Abdominal pain lower	2345	117	447	943	1898	5	7
Abdominal pain upper	11491	808	3595	2273	7896	22	57
Abdominal rebound tenderness	1	1	1				
Abdominal rigidity	121	10	44	22	77		1
Abdominal symptom	42	3	16	10	26		
Abdominal tenderness	129	15	57	19	72	1	1
Abdominal wall cyst	1		1				
Abdominal wall disorder	1			1	1		

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

System Organ Class

Gastrointestinal disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Abdominal wall haematoma	11	2	9		2		
Abdominal wall haemorrhage	3		3				
Abdominal wall oedema	4	1	2	1	2		
Abnormal faeces	858	25	88	211	770		
Acetonaemic vomiting	11	3	3	4	8		
Achlorhydria	2	1	1		1		
Acid peptic disease	1			1	1		
Acquired macroglossia	2		1		1		
Acquired oesophageal web	1				1		
Acute abdomen	39	7	39			1	1
Acute haemorrhagic ulcerative colitis	5	1	5				
Aerophagia	20	1	6	4	14		
Alcoholic pancreatitis	1		1				
Allergic gastroenteritis	6		2	1	4		
Allergic stomatitis	1				1		
Anaesthesia oral	109	2	18	23	91		
Anal blister	1				1		
Anal eczema	4	1	1	3	3		
Anal erosion	2			1	2		
Anal erythema	3		2		1		
Anal fissure	17		5	6	12		
Anal fistula	7	2	3	1	4		
Anal haemorrhage	83	9	35	20	48		
Anal hypoaesthesia	5	2	4		1		
Anal incontinence	194	20	94	26	100		1
Anal inflammation	5		1	2	4		
Anal paraesthesia	2	2	2				

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

System Organ Class

Gastrointestinal disorders

Preferred Term	Total # of Spontaneous AE	Spontaneous				Non Interventional Study	
		Serious		Nonserious		Serious	
		I	C	I	C	I	C
Anal pruritus	28	3	6	10	22		
Anal rash	4	1	2		2		
Anal skin tags	1				1		
Anal spasm	2			1	2		
Anal sphincter atony	27	4	15	4	12		
Anal sphincter hypertonia	1				1		
Anal ulcer	4	2	4				
Anastomotic ulcer perforation	1		1				
Angina bullosa haemorrhagica	8	2	8				
Angular cheilitis	36	2	5	10	31		
Anorectal discomfort	31	3	7	5	24		
Anorectal disorder	23	3	17	5	6		
Anorectal swelling	4		2	2	2		
Aphthous ulcer	915	33	136	213	779	1	5
Apical granuloma	2			1	2		
Appendicitis noninfective	1	1	1				1
Appendicolith	2	1	1		1		
Appendix disorder	16	3	13	1	3	2	3
Aptyalism	75	8	26	19	49		
Ascites	156	61	156				
Atrophic glossitis	5		1	2	4		
Autoimmune colitis	6		6				
Autoimmune enteropathy	1		1				
Autoimmune pancreatitis	18	7	18				
Barrett's oesophagus	9	4	9				1
Bezoar	1				1		
Bile acid malabsorption	9	4	6	1	3		

090177e19b0e6be1\Approved\Approved On: 19-Aug-2022 10:22 (GMT)

* I=Interval, C=Cumulative
 * AE=Adverse Event
 * Multiple adverse events coding to the same Preferred Term may have been reported in some cases. Counts are provided for the number of unique Preferred Terms reported.

