

ALLERGY/IMMUNOLOGY

Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Allergic reaction/hypersensitivity (including drug fever)</p> <p>Grade 1: transient rash, drug fever <38 degrees C (<100.4 degrees F) Grade 2: urticaria, drug fever >= 38 degrees C (>= 100.4 degrees F), and/or asymptomatic bronchospasm Grade 3: symptomatic bronchospasm, requiring parenteral medication(s), with or without urticaria; allergy-related edema/angioedema Grade 4: anaphylaxis Grade 5: death related to toxicity</p> <p>Note: Isolated urticaria, in the absence of other manifestations of an allergic or hypersensitivity reaction, is graded in the DERMATOLOGY/SKIN category.</p>	Allergic reaction	1, 2, 3, or 4	LLT : Allergic reaction	PT : Hypersensitivity Nos	HLT : Allergic conditions NEC	HLGT : Allergic conditions	Pr : Immune system disorders	SSC₁ : HYPERSENSITIVITY REACTIONS		
	Hypersensitivity	1, 2, 3, or 4	LLT : Hypersensitivity							
	Drug fever	1 (drug fever <38°C) 2 (drug fever > or = 38°C)	LLT : Drug fever	PT : Pyrexia	HLT : Febrile disorders	HLGT : Body temperature conditions	Pr : General disorders and administration site conditions			
	Rash	1	LLT : Rash	PT : Rash NOS	HLT : Rashes, eruptions and exanthems NEC	HLGT : Epidermal and dermal conditions	Pr : Skin and subcutaneous tissue disorders			
	Allergic rash	1	<p>If you need to set a complete mapping version (CTC versions 2.0 and/or 3.0 - MedDRA), please contact Pharmadhoc 112, rue Olivier de Serres 75015 Paris Tel: +33 (0)8 71 76 88 17 Tel: +33 (0)6 73 51 37 96 or send an email to info@pharmadhoc.com</p>							
	Hypersensitivity: rash	1	<p>If you need to set a complete mapping version (CTC versions 2.0 and/or 3.0 - MedDRA), please contact Pharmadhoc 112, rue Olivier de Serres 75015 Paris Tel: +33 (0)8 71 76 88 17 Tel: +33 (0)6 73 51 37 96 or send an email to info@pharmadhoc.com</p>							
	Urticaria	2	LLT : Urticaria	PT : Urticaria NOS	HLT : Urticarias	HLGT : Allergic conditions	Pr : Skin and subcutaneous tissue disorders	SSC₁ : ANAPHYLAXIS		
	Allergic urticaria	2	LLT : Allergic urticaria				Pr : Immune system disorders			
	Rash urticarial	2	LLT : Rash urticarial							
	Asymptomatic Bronchospasm	2	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :	SSC₂ :	
	Symptomatic Bronchospasm	3								
Allergy-related edema	3	LLT : Allergic edema	PT : Allergic oedema NOS	HLT : Allergic conditions NEC HLT : Oedema NEC	HLGT : Allergic conditions HLGT : General system disorders NEC	Pr : Immune system disorders Sec₁ : General disorders and administration site conditions	SSC₁ : OEDEMA			

Category: ALLERGY/IMMUNOLOGY

	Allergy-related angioedema	3	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :	SSC₂ :	SSC₃ :
						HLGT :	Sec₁ :			
<p>Adverse Event Allergic rhinitis (including sneezing, nasal stuffiness, postnasal drip)</p> <p>Grade 1 : mild, not requiring treatment</p> <p>Grade 2 : moderate, requiring treatment</p> <p>N/A Grade 3 :</p> <p>N/A Grade 4 :</p> <p>N/A Grade 5 :</p>	Allergic rhinitis	1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
					HLT :	HLGT :	Sec₁ :			
	Sneezing	1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
	Nasal stuffiness	1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
	Postnasal drip	1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Autoimmune reaction</p> <p>Grade 1 : serologic or other evidence of autoimmune reaction but patient is asymptomatic (e.g., vitiligo), all organ function is normal and no treatment is required</p> <p>Grade 2 : evidence of autoimmune reaction involving a non-essential organ or function (e.g., hypothyroidism), requiring treatment other than immunosuppressive drugs</p> <p>Grade 3 : reversible autoimmune reaction involving function of a major organ or other toxicity (e.g., transient colitis or anemia), requiring short-term immunosuppressive treatment</p> <p>Grade 4 : autoimmune reaction causing major grade 4 organ dysfunction; progressive and irreversible reaction; long-term administration of high-dose immunosuppressive therapy required</p> <p>Grade 5 : death related to toxicity</p> <p>Note: Also consider Hypothyroidism, Colitis, Hemoglobin, Hemolysis.</p>	Autoimmune reaction	1, 2, 3, or 4	LLT : Autoimmune disorder NOS	PT : Autoimmune disorder NOS	HLT : Autoimmune disorders NEC	HLGT : Autoimmune disorders	Pr : Immune system disorders			
<p>Adverse Event Serum sickness</p> <p>N/A Grade 1 :</p> <p>N/A Grade 2 :</p> <p>Grade 3 : present</p> <p>N/A Grade 4 :</p> <p>Grade 5 : death related to toxicity</p>	Serum sickness	3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Urticaria is graded in the DERMATOLOGY/SKIN category if it occurs as an isolated symptom. If it occurs with other manifestations of allergic or hypersensitivity reaction, grade as Allergic reaction/hypersensitivity above</p>										

<p><u>Adverse Event</u> Vasculitis</p> <p><u>Grade 1</u>: mild, not requiring treatment <u>Grade 2</u>: symptomatic, requiring medication <u>Grade 3</u>: requiring steroids <u>Grade 4</u>: ischemic changes or requiring amputation <u>Grade 5</u>: death related to toxicity</p>	Vasculitis	1, 2, 3, or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p><u>Adverse Event</u> Allergy-Other (Specify, _____)</p> <p><u>Grade 1</u>: mild <u>Grade 2</u>: moderate <u>Grade 3</u>: severe <u>Grade 4</u>: life-threatening or disabling <u>Grade 5</u>: death related to toxicity</p>					<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₁</u> :			
AUDITORY/HEARING										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3
Conductive hearing loss is graded as Middle ear/hearing in the AUDITORY/HEARING category										
Earache is graded in the PAIN category										
<p><u>Adverse Event</u> External Auditory Canal</p> <p><u>Grade 1</u>: external otitis with erythema or dry desquamation <u>Grade 2</u>: external otitis with moist desquamation <u>Grade 3</u>: external otitis with discharge, mastoiditis <u>Grade 4</u>: necrosis of the canal soft tissue or bone <u>Grade 5</u>: death related to toxicity</p> <p>Note: Changes associated with radiation to external ear (pinnae) are graded under Radiation dermatitis in the DERMATOLOGY/SKIN category.</p>	External Auditory Canal	1, 2, 3 or 4	<u>LLT</u> : Otitis externa (excl boil of meatus)	<u>PT</u> : Otitis externa NOS	<u>HLT</u> : Ear infections <u>HLT</u> : External ear infections and inflammations	<u>HLGT</u> : Infections - pathogen class unspecified <u>HLGT</u> : External ear disorders (excl congenital)	<u>Pr</u> : Infections and infestations <u>Sec₁</u> : Ear and labyrinth disorders			
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	External otitis	1, 2 or 3								
	Mastoiditis	3	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> : <u>HLT</u> :	<u>HLGT</u> : <u>HLGT</u> :	<u>Pr</u> : <u>Sec₁</u> :			
	Inner ear/hearing	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			

BLOOD/BONE MARROW											
Toxicity category and code name		Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
Category: AUDITORY/HEARING Adverse Event Inner ear/hearing Grade 1: hearing loss on audiometry only Grade 2: tinnitus or hearing loss, not requiring hearing aid or treatment Grade 3: tinnitus or hearing loss, correctable with hearing aid or treatment Grade 4: severe unilateral or bilateral hearing loss (deafness), not correctable Grade 5: death related to toxicity Adverse Event Middle ear/hearing Grade 1: serous otitis without subjective decrease in hearing Grade 2: serous otitis or infection requiring medical intervention; subjective decrease in hearing; rupture of tympanic membrane with discharge Grade 3: otitis with discharge, mastoiditis or conductive hearing loss Grade 4: necrosis of the canal soft tissue or bone Grade 5: death related to toxicity Adverse Event Auditory/Hearing-Other (Specify,_____) Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity		Inner ear	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		Hearing	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		Hearing loss	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		Tinnitus	2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec ₁ :			
		Deafness	4	LLT :	PT :	HLT :	HLGT :	Pr :			
		Middle ear/hearing	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		Serous otitis	1, 2 or 3			HLT :	HLGT :	Sec ₁ :			
		Rupture of tympanic membrane	2	LLT :	PT :	HLT :	HLGT :	Pr :			
		Mastoiditis	3	LLT :	PT :	HLT :	HLGT :	Pr :			
				HLT :	HLGT :	Sec ₁ :					
Conductive hearing loss	3	LLT : Conductive hearing loss	PT : Conductive deafness	HLT : Hearing losses	HLGT : Hearing disorders	Pr : Ear and labyrinth disorders					
BLOOD/BONE MARROW											
		Bone marrow cellularity	1, 2 or 3	LLT : Bone marrow hypocellular	PT : Bone marrow depression NOS	HLT : Marrow depression and hypoplastic anaemias	HLGT : Anaemias nonhaemolytic and marrow depression	Pr : Blood and lymphatic system disorders	SSC ₁ :	SSC ₂ :	
									BLOOD DYSCRASIAS/ BONE MARROW DEPRESSION	SECONDARY IMMUNOCOM PROMISED STATE	

<p>Grade 2: moderately hypocellular or >25 - <= 50% reduction from normal cellularity for age or >2 but <4 weeks to recovery of normal bone marrow cellularity</p> <p>Grade 3: severely hypocellular or >50 - <=75% reduction in cellularity for age or 4 - 6 weeks to recovery of normal bone marrow cellularity</p> <p>Grade 4: aplasia or >6 weeks to recovery of normal bone marrow cellularity</p> <p>Grade 5: death related to toxicity</p> <p>Note: Grade Bone marrow cellularity only for changes related to treatment, not disease. Normal ranges: children (<=18 years) 90% cellularity average; younger adults (19-59) 60-70% cellularity average; older adults (>60 years) 50% cellularity average.</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT : Haematological disorders</p>	<p>HLGT : Haematological disorders NEC</p>	<p>Pr : Blood and lymphatic system disorders</p>	<p>SSC₁ : BLOOD DYSCRASIAS/ BONE MARROW DEPRESSION</p>		
<p>Adverse Event CD4 count</p> <p>Grade 1: < LLN - 500/mm(3)</p> <p>Grade 2: 200 - <500/mm(3)</p> <p>Grade 3: 50 - <200/mm(3)</p> <p>Grade 4: <50/mm(3)</p> <p>Grade 5: death related to toxicity</p>	<p>CD4 count</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLGT : Immunology and allergy investigations</p>	<p>Pr : Investigations</p>			
<p>Adverse Event Haptoglobin</p> <p>Grade 1: decreased</p> <p>N/A Grade 2:</p> <p>Grade 3: absent</p> <p>N/A Grade 4:</p> <p>Grade 5: death related to toxicity</p>	<p>Haptoglobin</p>	<p>1 or 3</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT : Haem metabolism analyses</p>	<p>HLGT : Metabolic, nutritional and blood gas investigations</p>	<p>Pr : Investigations</p>		
<p>Adverse Event Hemoglobin</p> <p>Grade 1: < LLN - 10.0 g/dL; < LLN - 100 g/L; < LLN - 6.2 mmol/L</p> <p>Grade 2: 8.0 - <10.0 g/dL; 80 - <100 g/L; 4.9 - <6.2 mmol/L</p> <p>Grade 3: 6.5 - <8.0 g/dL; 65 - <80 g/L; 4.0 - <4.9 mmol/L</p> <p>Grade 4: <6.5 g/dL; <65 g/L; <4.0 mmol/L</p> <p>Grade 5: death related to toxicity</p>	<p>Hemoglobin</p>	<p>1, 2, 3 or 4</p>	<p>LLT : Hemoglobin decreased</p>	<p>PT : Haemoglobin decreased</p>	<p>HLT : Red blood cell analyses</p>	<p>HLGT : Haematology investigations (incl blood groups)</p>	<p>Pr : Investigations</p>		
<p>Adverse Event Hemoglobin for leukemia studies or bone marrow infiltrative/ myelophthisic processes, if specified in the protocol.</p> <p>Grade 1: 10 - <25% decrease from pretreatment</p> <p>Grade 2: 25 - <50% decrease from pretreatment</p> <p>Grade 3: 50 - <75% decrease from pretreatment</p> <p>Grade 4: >= 75% decrease from pretreatment</p> <p>Grade 5: death related to toxicity</p> <p>Note: The above criteria may be used for leukemia studies or bone marrow infiltrative/myelophthisic process if the protocol so specifies.</p>	<p>Hemoglobin</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>		

<p>Adverse Event Hemolysis (e.g., immune hemolytic anemia, drug related hemolysis, other)</p> <p>Grade 1: only laboratory evidence of hemolysis [e.g., direct antiglobulin test (DAT, Coombs) schistocytes]</p> <p>Grade 2: evidence of red cell destruction and ≥ 2gm decrease in hemoglobin, no transfusion</p> <p>Grade 3: requiring transfusion and/or medical intervention (e.g., steroids)</p> <p>Grade 4: catastrophic consequences of hemolysis (e.g., renal failure, hypotension, bronchospasm, emergency splenectomy)</p> <p>Grade 5: death related to toxicity</p> <p>Note: Also consider Haptoglobin, Hemoglobin.</p>	Hemolysis	1, 2, 3 or 4	LLT: Hemolysis	PT: Haemolysis NOS	HLT: Haemolyses NEC	HLGT: Haemolyses and related conditions	Pr: Blood and lymphatic system disorders			
<p>Adverse Event Leukocytes (total WBC)</p> <p>Grade 1: $< LLN - 3.0 \times 10^9/L$; $< LLN - 3000/mm(3)$</p> <p>Grade 2: $\geq 2.0 - <3.0 \times 10^9/L$; $\geq 2000 - <3000/mm(3)$</p> <p>Grade 3: $\geq 1.0 - <2.0 \times 10^9/L$; $\geq 1000 - <2000/mm(3)$</p> <p>Grade 4: $<1.0 \times 10^9/L$; $<1000/mm(3)$</p> <p>Grade 5: death related to toxicity</p>	Leukocytes	1, 2, 3 or 4	LLT:	PT:	HLT:	HLGT:	Pr:	SSC: BLOOD DYSCRASIAS/ BONE MARROW DEPRESSION		
<p>Adverse Event Leukocytes (total WBC) for BMT studies, if specified in the protocol.</p> <p>Grade 1: $\geq 2.0 - <3.0 \times 10^9/L$; $\geq 2000 - <3000/mm(3)$</p> <p>Grade 2: $\geq 1.0 - <2.0 \times 10^9/L$; $\geq 1000 - <2000/mm(3)$</p> <p>Grade 3: $\geq 0.5 - <1.0 \times 10^9/L$; $\geq 500 - <1000/mm(3)$</p> <p>Grade 4: $<0.5 \times 10^9/L$; $<500/mm(3)$</p> <p>Grade 5: death related to toxicity</p>	Leukocytes	1, 2, 3 or 4	LLT: Leukocyte count decreased	PT: White blood cell count decreased	HLT: White blood cell analyses	HLGT: Haematology investigations (incl blood groups)	Pr: Investigations	SSC: BLOOD DYSCRASIAS/ BONE MARROW DEPRESSION		
<p>Adverse Event Leukocytes (total WBC) for pediatric BMT studies (using age, race and sex normal values), if specified in the protocol.</p> <p>Grade 1: $\geq 75 - <100\%$ LLN</p> <p>Grade 2: $\geq 50 - <75\%$ LLN</p> <p>Grade 3: $\geq 25 - 50\%$ LLN</p> <p>Grade 4: $<25\%$ LLN</p> <p>Grade 5: death related to toxicity</p>	Leukocytes	1, 2, 3 or 4	LLT: Leukocyte count decreased	PT: White blood cell count decreased	HLT: White blood cell analyses	HLGT: Haematology investigations (incl blood groups)	Pr: Investigations	SSC: BLOOD DYSCRASIAS/ BONE MARROW DEPRESSION		
<p>Adverse Event Lymphopenia</p> <p>Grade 1: $< LLN - 1.0 \times 10^9/L$; $< LLN - 1000/mm(3)$</p> <p>Grade 2: $\geq 0.5 - <1.0 \times 10^9/L$; $\geq 500 - <1000/mm(3)$</p> <p>Grade 3: $<0.5 \times 10^9/L$; $<500/mm(3)$</p> <p>N/A Grade 4:</p> <p>Grade 5: death related to toxicity</p>	Lymphopenia	1, 2 or 3	LLT:	PT:	HLT:	HLGT: White blood cell disorders	Pr: Blood and lymphatic system disorders	SSC: BLOOD DYSCRASIAS/ BONE MARROW DEPRESSION		

Category : BLOOD/BONE MARROW

<p>Adverse Event Lymphopenia for pediatric BMT studies (using age, race and sex normal values), if specified in the protocol.</p> <p>Grade 1: ≥ 75 - $< 100\%$ LLN Grade 2: ≥ 50 - $< 75\%$ LLN Grade 3: ≥ 25 - $< 50\%$ LLN Grade 4: $< 25\%$ LLN Grade 5: Death due to toxicity</p>	Lymphopenia	1, 2, 3 or 4	LLT: Lymphopenia	PT: Lymphopenia	HLT: Leukopenias NEC	HLGT: White blood cell disorders	Pr: Blood and lymphatic system disorders	SSC: BLOOD DYSCRASIAS/ BONE MARROW DEPRESSION		
<p>Adverse Event Neutrophils/granulocytes (ANC/AGC)</p> <p>Grade 1: ≥ 1.5 - $< 2.0 \times 10^9/L$; ≥ 1500 - $< 2000/mm(3)$ Grade 2: ≥ 1.0 - $< 1.5 \times 10^9/L$; ≥ 1000 - $< 1500/mm(3)$ Grade 3: ≥ 0.5 - $< 1.0 \times 10^9/L$; ≥ 500 - $< 1000/mm(3)$ Grade 4: $< 0.5 \times 10^9/L$; $< 500/mm(3)$ Grade 5: death related to toxicity</p>	Neutrophils	1, 2, 3 or 4	LLT:	PT:	HLT:	HLGT:	Pr:	SSC:		
<p>Grade 1: ≥ 1.5 - $< 2.0 \times 10^9/L$; ≥ 1500 - $< 2000/mm(3)$ Grade 2: ≥ 1.0 - $< 1.5 \times 10^9/L$; ≥ 1000 - $< 1500/mm(3)$ Grade 3: ≥ 0.5 - $< 1.0 \times 10^9/L$; ≥ 500 - $< 1000/mm(3)$ Grade 4: $< 0.5 \times 10^9/L$; $< 500/mm(3)$ Grade 5: death related to toxicity</p>	Granulocytes	1, 2, 3 or 4	LLT: Granulocyte count decreased							
<p>Adverse Event Neutrophils/granulocytes (ANC/AGC) for BMT studies, if specified in the protocol.</p> <p>Grade 1: ≥ 1.0 - $< 1.5 \times 10^9/L$; ≥ 1000 - $< 1500/mm(3)$ Grade 2: ≥ 0.5 - $< 1.0 \times 10^9/L$; ≥ 500 - $< 1000/mm(3)$ Grade 3: ≥ 0.1 - $< 0.5 \times 10^9/L$; ≥ 100 - $< 500/mm(3)$ Grade 4: $< 0.1 \times 10^9/L$; $< 100/mm(3)$ Grade 5: death related to toxicity</p>	Neutrophils	1, 2, 3 or 4	LLT:	PT:	HLT:	HLGT:	Pr:	SSC:		
<p>Grade 1: ≥ 1.0 - $< 1.5 \times 10^9/L$; ≥ 1000 - $< 1500/mm(3)$ Grade 2: ≥ 0.5 - $< 1.0 \times 10^9/L$; ≥ 500 - $< 1000/mm(3)$ Grade 3: ≥ 0.1 - $< 0.5 \times 10^9/L$; ≥ 100 - $< 500/mm(3)$ Grade 4: $< 0.1 \times 10^9/L$; $< 100/mm(3)$ Grade 5: death related to toxicity</p>	Granulocytes	1, 2, 3 or 4	LLT:							
<p>Adverse Event Neutrophils/granulocytes (ANC/AGC) for leukemia studies or bone marrow infiltrative/myelophthisic process, if specified in the protocol.</p> <p>Grade 1: 10 - $< 25\%$ decrease from baseline Grade 2: 25 - $< 50\%$ decrease from baseline Grade 3: 50 - $< 75\%$ decrease from baseline Grade 4: $\geq 75\%$ decrease from baseline Grade 5: death related to toxicity</p>	Neutrophils	1, 2, 3 or 4	LLT:	PT:	HLT:	HLGT:	Pr:	SSC:		
<p>Grade 1: 10 - $< 25\%$ decrease from baseline Grade 2: 25 - $< 50\%$ decrease from baseline Grade 3: 50 - $< 75\%$ decrease from baseline Grade 4: $\geq 75\%$ decrease from baseline Grade 5: death related to toxicity</p>	Granulocytes	1, 2, 3 or 4	LLT:							
<p>Adverse Event Platelets</p> <p>Grade 1: $< LLN$ - $< 75.0 \times 10^9/L$; $< LLN$ - $75,000/mm(3)$ Grade 2: ≥ 50.0 - $< 75.0 \times 10^9/L$; $\geq 50,000$ - $< 75,000/mm(3)$ Grade 3: ≥ 10.0 - $< 50.0 \times 10^9/L$; $\geq 10,000$ - $< 50,000/mm(3)$ Grade 4: $< 10.0 \times 10^9/L$; $< 10,000/mm(3)$ Grade 5: death related to toxicity</p>	Platelets	1, 2, 3 or 4	LLT:	PT:	HLT:	HLGT:	Pr:	SSC:		
<p>Adverse Event Platelets for BMT studies, if specified in the protocol.</p> <p>Grade 1: ≥ 50.0 - $< 75.0 \times 10^9/L$; $\geq 50,000$ - $< 75,000/mm(3)$ Grade 2: ≥ 20.0 - $< 50.0 \times 10^9/L$; $\geq 20,000$ - $< 50,000/mm(3)$ Grade 3: ≥ 10.0 - $< 20.0 \times 10^9/L$; $\geq 10,000$ - $< 20,000/mm(3)$ Grade 4: $< 10.0 \times 10^9/L$; $< 10,000/mm(3)$ Grade 5: death related to toxicity</p>	Platelets	1, 2, 3 or 4	LLT:	PT:	HLT:	HLGT:	Pr:	SSC:		

<p>Adverse Event Platelets for leukemia studies or bone marrow infiltrative/myelophthistic process, if specified in the protocol.</p> <p>Grade 1: 10 - <25% decrease from baseline Grade 2: 25 - <50% decrease from baseline Grade 3: 50 - <75% decrease from baseline Grade 4: >= 75% decrease from baseline Grade 5: death related to toxicity</p>	Platelets	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
<p>Adverse Event Transfusion: Platelets</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: yes Grade 4: platelet transfusions and other measures required to improve platelet increment; platelet transfusion refractoriness associated with life-threatening bleeding. (e.g., HLA or cross matched platelet transfusions) Grade 5: death related to toxicity</p> <p>Note: Also consider Platelets</p>	Platelets transfusion	3 or 4	LLT : Platelets transfusion	PT : Platelets transfusion	HLT : Blood and blood product treatment	HLGT : Haematological and lymphoid tissue therapeutic procedures	Pr : Surgical and medical procedures			
<p>Adverse Event Transfusion: Platelets for BMT studies, if specified in the protocol.</p> <p>Grade 1: 1 platelet transfusion in 24 hours Grade 2: 2 platelet transfusions in 24 hours Grade 3: >= 3 platelet transfusions in 24 hours Grade 4: platelet transfusions and other measures required to improve platelet increment; platelet transfusion refractoriness associated with life-threatening bleeding. (e.g., HLA or cross matched platelet transfusions) Grade 5: death related to toxicity</p> <p>Note: Also consider Platelets</p>	Platelets transfusion	3 or 4	LLT : Platelets transfusion	PT : Platelets transfusion	HLT : Blood and blood product treatment	HLGT : Haematological and lymphoid tissue therapeutic procedures	Pr : Surgical and medical procedures			
<p>Adverse Event Transfusion: pRBCs</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: yes N/A Grade 4: Grade 5: death related to toxicity</p> <p>Note: Also consider Hemoglobin</p>		3	LLT :	PT :	HLT :	HLGT :	Pr : Surgical and medical procedures			
<p>Adverse Event Transfusion: pRBCs for BMT studies, if specified in the protocol.</p> <p>Grade 1: <= 2 u pRBC in 24 hours elective or planned Grade 2: 3 u pRBC in 24 hours elective or planned Grade 3: >=4 u pRBC in 24 hours Grade 4: hemorrhage or hemolysis associated with life-threatening anemia; medical intervention required to improve hemoglobin Grade 5: death related to toxicity</p> <p>Note: Also consider Hemoglobin</p>	Transfusion pRBCs	1, 2, 3 or 4	LLT : Packed red blood cell transfusion	PT : Packed red blood cell transfusion	HLT : Blood and blood product treatment	HLGT : Haematological and lymphoid tissue therapeutic procedures	Pr : Surgical and medical procedures			

<p>Adverse Event Transfusion: pRBCs for pediatric BMT studies, if specified in the protocol.</p> <p>Grade 1: <=15mL/kg in 24 hours elective or planned Grade 2: >15 - <=30mL/kg in 24 hours elective or planned Grade 3: >30mL/kg in 24 hours Grade 4: hemorrhage or hemolysis associated with life-threatening anemia; medical intervention required to improve hemoglobin Grade 5: death related to toxicity</p> <p>Note: Also consider Hemoglobin</p>	<p>Transfusion pRBCs</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT : Blood and blood product treatment</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Blood/Bone Marrow-Other (Specify,_____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
CARDIOVASCULAR (ARRHYTHMIA)										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Conduction abnormality/Atrioventricular heart block</p> <p>Grade 1: asymptomatic, not requiring treatment (e.g., Mobitz type I second-degree AV block, Wenckebach) Grade 2: symptomatic, but not requiring treatment Grade 3: symptomatic and requiring treatment (e.g., Mobitz type II second-degree AV block, third-degree AV block) Grade 4: life-threatening (e.g., arrhythmia associated with CHF, hypotension, syncope, shock) Grade 5: death related to toxicity</p>	<p>Conduction abnormality/Atrioventricular heart block</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT : Cardiac arrhythmias</p>	<p>Pr : Cardiac disorders</p>	<p>SSC₁ : CARDIAC ISCHAEMIA</p>		
<p>Adverse Event Nodal/junctional arrhythmia/dysrhythmia</p> <p>Grade 1: asymptomatic, not requiring treatment Grade 2: symptomatic, but not requiring treatment Grade 3: symptomatic and requiring treatment Grade 4: life-threatening (e.g., arrhythmia associated with CHF, hypotension, syncope, shock) Grade 5: death related to toxicity</p>	<p>Conduction abnormality</p>	<p>1, 2, 3 or 4</p>	<p>If you need to set a complete mapping version (CTC versions 2.0 and/or 3.0 - MedDRA), please contact Pharmadhoc 112, rue Olivier de Serres 75015 Paris Tel: +33 (0)8 71 76 88 17 Tel: +33 (0)6 73 51 37 96 or send an email to info@pharmadhoc.com</p>							
		<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr : Cardiac disorders</p>			
		<p>1, 2, 3 or 4</p>	<p>LLT :</p>			<p>HLGT :</p>				
	<p>Dysrhythmia</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
	<p>Arrhythmia</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			

Category: CARDIOVASCULAR (ARRHYTHMIA)

<p>Adverse Event Palpitations</p> <p>Grade 1: present N/A Grade 2: N/A Grade 3: N/A Grade 4: N/A Grade 5:</p> <p>Note: Grade palpitations only in the absence of a documented arrhythmia.</p>		1	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Prolonged QTc interval (QTc > 0.48 seconds)</p> <p>Grade 1: asymptomatic, not requiring treatment Grade 2: symptomatic, but not requiring treatment Grade 3: symptomatic and requiring treatment Grade 4: life-threatening (e.g., arrhythmia associated with CHF, hypotension, syncope, shock) Grade 5: death related to toxicity</p>	Prolonged QTc interval (QTc > 0.48 seconds)		LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Sinus bradycardia</p> <p>Grade 1: asymptomatic, not requiring treatment Grade 2: symptomatic, but not requiring treatment Grade 3: symptomatic and requiring treatment Grade 4: life-threatening (e.g., arrhythmia associated with CHF, hypotension, syncope, shock) Grade 5: death related to toxicity</p>	Sinus bradycardia		LLT :	PT :	HLT :	HLGT :	Pr :			
	Bradycardia		LLT :	PT :	HLT : Rate and rhythm disorders NEC	HLGT :	Pr :			
<p>Adverse Event Sinus tachycardia</p> <p>Grade 1: asymptomatic, not requiring treatment Grade 2: symptomatic, but not requiring treatment Grade 3: symptomatic and requiring treatment of underlying cause N/A Grade 4: Grade 5: death related to toxicity</p>		1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Supraventricular arrhythmias (SVT/atrial fibrillation/flutter)</p> <p>Grade 1: asymptomatic, not requiring treatment Grade 2: symptomatic, but not requiring treatment Grade 3: symptomatic and requiring treatment Grade 4: life-threatening (e.g., arrhythmia associated with CHF, hypotension, syncope, shock) Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	SVT		LLT :							
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT : Cardiac arrhythmias	Pr :			

		1, 2, 3 or 4	LLT :	PT :	HLT : Supraventricular arrhythmias	HLGT : Cardiac arrhythmias	Pr : Cardiac disorders				
Syncope (fainting) is graded in the NEUROLOGY category											
<p>Adverse Event Vasovagal episode</p> <p>N/A Grade 1:</p> <p>Grade 2: present without loss of consciousness</p> <p>Grade 3: present with loss of consciousness</p> <p>N/A Grade 4:</p> <p>Grade 5: death related to toxicity</p>	Vasovagal episode	2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :				
		3	LLT :	PT :	HLT :	HLGT :	Pr :				
<p>Adverse Event Ventricular arrhythmia (PVCs/bigeminy/trigeminy/ventricular tachycardia)</p> <p>Grade 1: asymptomatic, not requiring treatment</p> <p>Grade 2: symptomatic, but not requiring treatment</p> <p>Grade 3: symptomatic and requiring treatment</p> <p>Grade 4: life-threatening (e.g., arrhythmia associated with CHF, hypotension, syncope, shock)</p> <p>Grade 5: death related to toxicity</p>	Ventricular arrhythmia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Cardiac disorders	SSC₁ : ARREST (CARDIAC)	SSC₂ : CARDIAC ISCHAEMIA		
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Cardiac disorders				
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Cardiac disorders				
	Trigeminy			LLT :	PT :	HLT : Ventricular arrhythmias and cardiac arrest	HLGT : Cardiac arrhythmias	Pr : Cardiac disorders			
		1, 2, 3 or 4	LLT :	PT :	HLT : HLT : Autonomic nervous system disorders	HLGT : HLGT :	Pr : Sec₁ : Nervous system disorders	SSC₁ : ARREST (CARDIAC)	SSC₂ : CARDIAC ISCHAEMIA		
<p>Adverse Event Cardiovascular/Arrhythmia-Other (Specify,_____)</p> <p>Grade 1: asymptomatic, not requiring treatment</p> <p>Grade 2: symptomatic, but not requiring treatment</p> <p>Grade 3: symptomatic, and requiring treatment of underlying cause</p> <p>Grade 4: life-threatening (e.g., arrhythmia associated with CHF, hypotension, syncope, shock)</p> <p>Grade 5: death related to toxicity</p>											
CARDIOVASCULAR (GENERAL)											
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3	

<p>Adverse Event Acute vascular leak syndrome</p> <p>N/A Grade 1: Grade 2: symptomatic, but not requiring fluid support Grade 3: respiratory compromise or requiring fluids Grade 4: life-threatening; requiring pressor support and/or ventilatory support Grade 5: death related to toxicity</p>	Acute vascular leak syndrome		LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Cardiac-ischemia/infarction</p> <p>Grade 1: non-specific T-wave flattening or changes Grade 2: asymptomatic, ST- and T- wave changes suggesting ischemia Grade 3: angina without evidence of infarction Grade 4: acute myocardial infarction Grade 5: death related to toxicity</p>	Cardiac-ischemia/infarction	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Cardiac disorders	SSC :		
			LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Cardiac left ventricular function</p> <p>Grade 1: asymptomatic decline of resting ejection fraction of $\geq 10\%$ but $< 20\%$ of baseline value; shortening fraction $\geq 24\%$ but $< 30\%$ Grade 2: asymptomatic but resting ejection fraction below LLN for laboratory or decline of resting ejection fraction $\geq 20\%$ of baseline value; $< 24\%$ shortening fraction Grade 3: CHF responsive to treatment Grade 4: severe or refractory CHF or requiring intubation Grade 5: death related to toxicity</p>	Cardiac left ventricular function		LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
CNS cerebrovascular ischemia is graded in the NEUROLOGY category										
<p>Adverse Event Cardiac troponin I (cTnI)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: levels consistent with unstable angina as defined by the manufacturer Grade 4: levels consistent with myocardial infarction as defined by the manufacturer Grade 5: death related to toxicity</p>	Cardiac troponin I (cTnI)	3 or 4	LLT :	PT :	HLT :	HLGT : Enzyme investigations NEC	Pr :			
<p>Adverse Event Cardiac troponin T (cTnT)</p> <p>Grade 1: ≥ 0.03 - < 0.05 ng/mL Grade 2: ≥ 0.05 - < 0.1 ng/mL Grade 3: ≥ 0.1 - < 0.2 ng/mL Grade 4: ≥ 0.2 ng/mL Grade 5: death related to toxicity</p>	Cardiac troponin T (cTnT)		LLT :	PT :	HLT :	HLGT :	Pr :			
					HLT :	HLGT :	Pr :			

ory: CARDIOVASCULAR (GENERAL)

<p>Adverse Event Edema</p> <p>Grade 1: asymptomatic, not requiring therapy Grade 2: symptomatic, requiring therapy Grade 3: symptomatic edema limiting function and unresponsive to therapy or requiring drug discontinuation Grade 4: anasarca (severe generalized edema) Grade 5: death related to toxicity</p>	Edema	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Sec₁ :	SSC₁ :	SSC₂ :		
					HLT :	HLGT :	Sec₂ :				
				LLT :	PT :	HLT :	HLGT :	Pr :			
				LLT :	PT :	HLT :	HLGT :	Pr :			
				LLT :	PT :	HLT : Heart failure signs and symptoms	HLGT :	Sec₁ :	SSC₁ :	SSC₂ :	
						HLT :	HLGT :	Sec₂ :			
<p>Adverse Event Hypertension</p> <p>Grade 1: asymptomatic, transient increase by >20 mmHg (diastolic) or to > 150/100* if previously WNL; not requiring treatment Grade 2: recurrent or persistent or symptomatic increase by > 20 mmHg (diastolic) or to > 150/100* if previously WNL; not requiring treatment Grade 3: requiring therapy or more intensive therapy than previously Grade 4: hypertensive crisis Grade 5: death related to toxicity</p> <p>Note: * For pediatric patients, use age and sex appropriate normal values >95th percentile ULN.</p>	Hypertension	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
			LLT :	PT :	HLT :	HLGT :	Pr :				
<p>Adverse Event Hypotension</p> <p>Grade 1: changes, but not requiring therapy (including transient orthostatic hypotension) Grade 2: requiring brief fluid replacement or other therapy but not hospitalization; no physiologic consequences Grade 3: requiring therapy and sustained medical attention, but resolves without persisting physiologic consequences Grade 4: shock (associated with acidemia and impairing vital organ function due to tissue hypoperfusion) Grade 5: death related to toxicity</p> <p>Note: Angina or MI is graded as Cardiac-ischemia/infarction in the CARDIOVASCULAR (GENERAL) category. For pediatric patients, systolic BP 65 mmHg or less in infants up to 1 year old and 70 mmHg or less in children older than 1 year of age, use two successive or three measurements in 24 hours. Also consider Syncope (fainting).</p>	Hypotension	1, 2, 3 or 4	LLT : Hypotension	PT : Hypotension NOS	HLT : Vascular hypotensive disorders	HLGT : Decreased and nonspecific blood pressure disorders and shock	Pr : Vascular disorders	SSC₁ : ANAPHYLAXIS	SSC₂ : CARDIAC ISCHAEMIA		
		1	LLT :	PT :	HLT :	HLGT :	Pr :				
	Hypovolemic shock		LLT :	PT :	HLT :	HLGT :	Pr :				
<p>Adverse Event Myocarditis</p> <p>N/A Grade 1: N/A Grade 2:</p>	Myocarditis	3 or 4	LLT : Myocarditis	PT : Myocarditis NOS	HLT : Noninfectious myocarditis	HLGT : Myocardial disorders	Pr : Cardiac disorders				

Categori	<p>Grade 3: CHF responsive to treatment Grade 4: severe or refractory CHF Grade 5: death related to toxicity</p>		3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :			
	<p>Adverse Event Operative injury of vein/artery</p> <p>Grade 1: primary suture repair for injury, but not requiring transfusion Grade 2: primary suture repair for injury, requiring transfusion Grade 3: vascular occlusion requiring surgery or bypass for injury Grade 4: myocardial infarction; resection of organ (e.g., bowel, limb) Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Injury, poisoning and procedural complications				
						HLT :	HLGT :	Sec ₁ : Vascular disorders				
			1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Injury, poisoning and procedural complications				
						HLT :	HLGT :	Sec ₁ : Vascular disorders				
			3	LLT :	PT :	HLT :	HLGT :	Pr :				
		Myocardial infarction	4	LLT : Myocardial infarction	PT : Myocardial infarction	HLT : Ischaemic coronary artery disorders	HLGT : Coronary artery disorders	Pr : Cardiac disorders				
	<p>Adverse Event Pericardial effusion/pericarditis</p> <p>Grade 1: asymptomatic effusion, not requiring treatment Grade 2: pericarditis (rub, ECG changes, and/or chest pain) Grade 3: with physiologic consequences Grade 4: tamponade (drainage or pericardial window required) Grade 5: death related to toxicity</p>	Pericardial effusion/pericarditis	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
				2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Sec ₁ : Infections and infestations			
	<p>Adverse Event Peripheral arterial ischemia</p> <p>N/A Grade 1: Grade 2: brief episode of ischemia managed non-surgically and without permanent deficit Grade 3: requiring surgical intervention Grade 4: life-threatening or with permanent functional deficit (e.g., amputation) Grade 5: death related to toxicity</p>	Peripheral arterial ischemia	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
<p>Adverse Event Phlebitis (superficial)</p> <p>N/A Grade 1: Grade 2: present N/A Grade 3: N/A Grade 4: N/A Grade 5:</p> <p>Note: Injection site reaction is graded in the DERMATOLOGY/SKIN category. Thrombosis/embolism is graded in the CARDIOVASCULAR (GENERAL) category.</p>	Phlebitis (superficial)		LLT :	PT :	HLT :	HLGT :	Pr :					

Syncope (fainting) is graded in the NEUROLOGY category										
<p>Adverse Event Thrombosis/embolism</p> <p>N/A Grade 1:</p> <p>Grade 2: deep vein thrombosis, not requiring anticoagulant</p> <p>Grade 3: deep vein thrombosis, requiring anticoagulant therapy</p> <p>Grade 4: embolic event including pulmonary embolism</p> <p>Grade 5: death related to toxicity</p>	Thrombosis/embolism	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :		
		2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
		4	LLT :	PT :	HLT :	HLGT :	Pr :			
		4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : THROMBOSIS		
Vein/artery operative injury is graded as Operative injury of vein/artery in the CARDIOVASCULAR (GENERAL) category										
<p>Adverse Event Visceral arterial ischemia (non-myocardial)</p> <p>N/A Grade 1:</p> <p>Grade 2: brief episode of ischemia managed non-surgically and without permanent deficit</p> <p>Grade 3: requiring surgical intervention</p> <p>Grade 4: life-threatening or with permanent functional deficit (e.g., resection of ileum)</p> <p>Grade 5: death related to toxicity</p>	Visceral arterial ischemia (non-myocardial)		LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Cardiovascular/General-Other (Specify,_____)</p> <p>Grade 1: mild</p> <p>Grade 2: moderate</p> <p>Grade 3: severe</p> <p>Grade 4: life-threatening or disabling</p> <p>Grade 5: death related to toxicity</p>										
COAGULATION										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3
Note: See the HEMORRHAGE category for grading the severity of bleeding events.										
<p>Adverse Event DIC (disseminated intravascular coagulation)</p> <p>N/A Grade 1:</p> <p>N/A Grade 2:</p> <p>Grade 3: laboratory findings present with no bleeding</p>	DIC	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			

Category: COAGULATION	<p><u>Grade 4</u>: laboratory findings and bleeding <u>Grade 5</u>: death related to toxicity</p> <p>Note: Must have increased fibrin split products or D-dimer in order to grade as DIC. Also consider Platelets.</p>			<u>LLT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Sec</u> :				
	<p><u>Adverse Event</u> Fibrinogen</p> <p><u>Grade 1</u>: ≥ 0.75 - < 1.0 x LLN <u>Grade 2</u>: ≥ 0.5 - < 0.75 x LLN <u>Grade 3</u>: ≥ 0.25 - < 0.5 x LLN <u>Grade 4</u>: < 0.25 x LLN <u>Grade 5</u>: death related to toxicity</p>	Fibrinogen	1, 2, 3 or 4	<p style="text-align: center;">If you need to set a complete mapping version (CTC versions 2.0 and/or 3.0 - MedDRA), please contact Pharmadhoc 112, rue Olivier de Serres 75015 Paris Tel: +33 (0)8 71 76 88 17 Tel: +33 (0)6 73 51 37 96 or send an email to info@pharmadhoc.com</p>							
	<p><u>Adverse Event</u> Fibrinogen for leukemia studies or bone marrow infiltrative/myelophthisic process, if specified in the protocol.</p> <p><u>Grade 1</u>: $< 20\%$ decrease from pretreatment value or LLN <u>Grade 2</u>: ≥ 20 - $< 40\%$ decrease from pretreatment value or LLN <u>Grade 3</u>: ≥ 40 - $< 70\%$ decrease from pretreatment value or LLN <u>Grade 4</u>: < 50 mg <u>Grade 5</u>: death related to toxicity</p>	Fibrinogen	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
	<p><u>Adverse Event</u> Partial thromboplastin time (PTT)</p> <p><u>Grade 1</u>: $> \text{ULN}$ - ≤ 1.5 x ULN <u>Grade 2</u>: > 1.5 - ≤ 2 x ULN <u>Grade 3</u>: > 2 x ULN <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>	Partial thromboplastin time (PTT)		<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
	Phelbitis is graded in the CARDIOVASCULAR (GENERAL) category										
	<p><u>Adverse Event</u> Prothrombin time (PT)</p> <p><u>Grade 1</u>: $> \text{ULN}$ - ≤ 1.5 x ULN <u>Grade 2</u>: > 1.5 - ≤ 2 x ULN <u>Grade 3</u>: > 2 x ULN <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>	Prothrombin time (PT)	1, 2 or 3	<u>LLT</u> : Prothrombin time prolonged	<u>PT</u> : Prothrombin time prolonged	<u>HLT</u> : Coagulation and bleeding analyses	<u>HLGT</u> : Haematology investigations (incl blood groups)	<u>Pr</u> : Investigations			
	Thrombosis/embolism is graded in the CARDIOVASCULAR (GENERAL) category										
			3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			

<p>Adverse Event Thrombotic microangiopathy (e.g., thrombotic thrombocytopenic purpura/TTP or hemolytic uremic syndrome/HUS)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: laboratory findings present without clinical consequences Grade 4: laboratory findings and clinical consequences, (e.g., CNS hemorrhage/bleeding or thrombosis/embolism or renal failure) requiring therapeutic intervention Grade 5: death related to toxicity</p> <p>Note: Must have microangiopathic changes on blood smear (e.g., schistocytes, helmet cells, red cell fragments). Also Consider Hemoglobin, Platelets, Creatinine.</p>		3 or 4	LLT :	PL :	HLT :	HLGT :	Sec₁ :			
					HLT :	HLGT :	Pr :			
		3 or 4	LLT :	PT :	HLT :	HLGT :	Sec₁ :	SSC₁ :		
			LLT :		HLT :	HLGT :	Sec₂ :			
	Hemolytic uremic syndrome	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :		HLT :	HLGT :	Sec₁ :			
<p>Adverse Event Thrombotic microangiopathy (e.g., thrombotic thrombocytopenic purpura/TTP or hemolytic uremic syndrome/HUS) for BMT studies, if specified by the protocol.</p> <p>Grade 1: evidence of RBC destruction (schistocytosis) without clinical consequences Grade 2: evidence of RBC destruction with elevated creatinine (<=3 x ULN) Grade 3: evidence of RBC destruction with creatinine (>3 x ULN) not requiring dialysis Grade 4: evidence of RBC destruction with renal failure requiring dialysis and/or encephalopathy Grade 5: death related to toxicity</p> <p>Note: Must have microangiopathic changes on blood smear (e.g., schistocytes, helmet cells, red cell fragments). Also Consider Hemoglobin, Platelets, Creatinine.</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Blood and lymphatic system disorders			
					HLT :	HLGT :	Sec₁ : Renal and urinary disorders			
		3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :		
			LLT :		HLT :	HLGT :	Sec₁ :	THROMBOSIS		
	Hemolytic uremic syndrome	3 or 4	LLT :		HLT :	HLGT : Renal disorders (excl nephropathies)	Pr : Renal and urinary disorders			
			LLT :		HLT :	HLGT : Haemolyses and related conditions	Sec₁ : Blood and lymphatic system disorders			
<p>Adverse Event Coagulation-Other (Specify,_____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
CONSTITUTIONAL SYMPTOMS										

Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Fatigue (lethargy, malaise, asthenia)</p> <p>Grade 1: increased fatigue over baseline, but not altering normal activities Grade 2: moderate (e.g., decrease in performance status by 1 ECOG level or 20% Karnofsky or Lansky) or causing difficulty performing some activities Grade 3: severe (e.g., decrease in performance status by ≥ 2 ECOG levels or 40% Karnofsky or Lansky) or loss of ability to perform some activities Grade 4: bedridden or disabling Grade 5: death related to toxicity</p> <p>Note: See Appendix III for performance status scales.</p>	Fatigue (lethargy, malaise, asthenia)	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Lethargy		LLT :	PT :	HLT :	HLGT :	Pr :			
	Malaise		LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Fever (in the absence of neutropenia, where neutropenia is defined as AGC<1.0 x 10e9/L)</p> <p>Grade 1: 38.0 - 39.0 C (100.4 - 102.2 F) Grade 2: 39.1 - 40.0 C (102.3 - 104.0 F) Grade 3: >40.0 C (>104.0 F) for <24hrs Grade 4: >40.0 C (>104.0 F) for > 24hrs Grade 5: death related to toxicity</p> <p>Note: The temperature measurements listed above are oral or tympanic. Also consider Allergic reaction/hypersensitivity.</p>	Fever	1, 2, 3 or 4	LLT : Fever	PT : Pyrexia	HLT : Febrile disorders	HLGT : Body temperature conditions	Pr : General disorders and administration site conditions			
<p>Hot flashes/flushes are graded in the ENDOCRINE category</p>										
<p>Adverse Event Rigors, chills</p> <p>Grade 1: mild, requiring symptomatic treatment (e.g., blanket) or non-narcotic medication Grade 2: severe and/or prolonged, requiring narcotic medication Grade 3: not responsive to narcotic medication N/A Grade 4: Grade 5: death related to toxicity</p>	Rigors, chills	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :							
<p>Adverse Event Sweating (diaphoresis)</p> <p>Grade 1: mild and occasional Grade 2: frequent or drenching N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Sweating (diaphoresis)		LLT :							
		1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
		1 or 2	LLT :							

pty: CONSTITUTIONAL SYMPTOMS

Categc	<p>Adverse Event Weight gain</p> <p>Grade 1: 5 - <10% Grade 2: 10 - <20% Grade 3: >=20% N/A Grade 4: Grade 5: death related to toxicity</p>	Weight gain		LLT :	PT :	HLT :	HLGT :	Pr :			
	<p>Adverse Event Weight gain - Veno-Occlusive Disease (VOD) for BMT studies if specified in the protocol.</p> <p>Grade 1: >=2 - <5% Grade 2: >=5 - <10% Grade 3: >=10% or as ascites Grade 4: >=10% or fluid retention resulting in pulmonary failure Grade 5: death related to toxicity</p> <p>Note: Also consider Ascites, Edema, Pleural effusion (non-malignant)</p>	Weight gain		LLT :	PT :	HLT :	HLGT :	Pr :			
			1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	<p>Adverse Event Weight loss</p> <p>Grade 1: 5 - <10% Grade 2: 10 - <20% Grade 3: >=20% N/A Grade 4: Grade 5: death related to toxicity</p>	Weight loss	1, 2, 3 or 4	LLT : Weight loss	PT : Weight decreased	HLT : Physical examination procedures	HLGT : Physical examination topics	Pr : Investigations			
	<p>Adverse Event Constitutional Symptoms-Other (Specify, _____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
DERMATOLOGY/SKIN											
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3	
<p>Adverse Event Alopecia</p> <p>Grade 1: mild hair loss Grade 2: pronounced hair loss N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Alopecia	1 or 2	LLT : Alopecia	PT : Alopecia	HLT : Alopecias	HLGT : Skin appendage conditions	Pr : Skin and subcutaneous tissue disorders				

<p>Adverse Event Bruising (in absence of grade 3 or 4 thrombocytopenia)</p> <p>Grade 1: localized or in dependent area Grade 2: generalized N/A Grade 3: N/A Grade 4: N/A Grade 5:</p> <p>Note: Bruising resulting from grade 3 or 4 thrombocytopenia is graded as Petechiae/purpura and Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia in the HEMORRHAGE category, not in the DERMATOLOGY/SKIN category.</p>	<p>Bruising (in absence of grade 3 or 4 thrombocytopenia)</p>		<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Dermatitis, focal (associated with high-dose chemotherapy and bone marrow transplant)</p> <p>Grade 1: faint erythema or dry desquamation Grade 2: moderate to brisk erythema or a patchy moist desquamation, mostly confined to skin folds and creases; moderate edema Grade 3: confluent moist desquamation, >1.5 cm diameter, not confined to skin folds; pitting edema Grade 4: Iskin necrosis or ulceration of full thickness dermis; may include spontaneous bleeding not induced by minor trauma or abrasion</p>			<p>If you need to set a complete mapping version (CTC versions 2.0 and/or 3.0 - MedDRA), please contact Pharmadhoc 112, rue Olivier de Serres 75015 Paris Tel: +33 (0)8 71 76 88 17 Tel: +33 (0)6 73 51 37 96 or send an email to info@pharmadhoc.com</p>							
<p>Adverse Event Dry skin</p> <p>Grade 1: controlled with emollients Grade 2: not controlled with emollients N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	<p>Dry skin</p>		<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Erythema multiforme (e.g., Stevens-Johnson syndrome, toxic epidermal necrolysis)</p> <p>N/A Grade 1: Grade 2: scattered, but not generalized eruption Grade 3: severe or requiring IV fluids (e.g., generalized rash or painful stomatitis) Grade 4: life-threatening (e.g., exfoliative or ulcerating dermatitis or requiring enteral or parenteral nutritional support) Grade 5: death related to toxicity</p>	<p>Erythema multiforme (e.g., Stevens-Johnson syndrome, toxic epidermal necrolysis)</p>		<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
		<p>2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
	<p>Toxic epidermal necrolysis</p>		<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>	<p>Sec₁ : Infections and infestations</p>		

<p>Adverse Event Flushing</p> <p>Grade 1: present N/A Grade 2: N/A Grade 3: N/A Grade 4: N/A Grade 5:</p> <p>Attribution Code Unrelated Unlikely Possible Probable Definite</p>	Flushing		LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hand-foot skin reaction</p> <p>Grade 1: skin changes or dermatitis without pain (e.g., erythema, peeling) Grade 2: skin changes with pain, not interfering with function Grade 3: skin changes with pain, interfering with function N/A Grade 4: Grade 5: death related to toxicity</p>	Hand-foot skin reaction	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Injection site reaction</p> <p>Grade 1: pain or itching or erythema Grade 2: pain or swelling, with inflammation or phlebitis Grade 3: ulceration or necrosis that is severe or prolonged, or requiring surgery N/A Grade 4: Grade 5: death related to toxicity</p>	Injection site reaction		LLT :	PT :	HLT :	HLGT :	Pr :			
	Injection site pain		LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : PAIN		
	Injection site itching		LLT :	PT :	HLT :	HLGT :	Pr : General disorders and administration site conditions	Sec₁ : Injury, poisoning and procedural complications		
		1	LLT :	PT :	HLT :	HLGT :	Pr :	Sec₁ :		
	Injection site swelling		LLT :	PT :	HLT :	HLGT :	Pr :	Sec₁ :		
		2	LLT :	PT :	HLT :	HLGT :	Pr :			

TOLOGY/SKIN			LLT	PT	HLT	HLGT	Sec₁				
	Injection site phlebitis		LLT	PT	HLT	HLGT	Pr				
			LLT	PT	HLT	HLGT	Sec₂ : Vascular disorders				
			LLT	PT	HLT	HLGT	Pr				
	Injection site necrosis	3	LLT	PT	HLT	HLGT	Sec₁				
			LLT	PT	HLT	HLGT	Pr				
	Adverse Event Nail changes Grade 1 : discoloration or ridging (koilonychia) or pitting Grade 2 : partial or complete loss of nail(s) or pain in nailbeds N/A Grade 3 : N/A Grade 4 : N/A Grade 5 :	Nail changes		LLT	PT	HLT	HLGT	Pr			
		Nail discolouration	1 or 2	LLT	PT	HLT	HLGT	Pr			
			1 or 2	LLT	PT	HLT	HLGT	Pr			
		Loss of nail(s)	2	LLT	PT	HLT	HLGT	Pr			
	Petechiae is graded in the HEMORRHAGE category.										
	Adverse Event Photosensitivity Grade 1 : painless erythema Grade 2 : painful erythema Grade 3 : erythema with desquamation N/A Grade 4 : Grade 5 : death related to toxicity	Photosensitivity		LLT	PT	HLT	HLGT	Pr			
	Adverse Event Pigmentation changes (e.g., vitiligo) Grade 1 : localized pigmentation changes Grade 2 : generalized pigmentation changes N/A Grade 3 : N/A Grade 4 : N/A Grade 5 :	Pigmentation changes (e.g., vitiligo)		LLT	PT	HLT	HLGT : Pigmentation disorders	Pr : Skin and subcutaneous tissue disorders			
		Vitiligo		LLT	PT	HLT	HLGT : Pigmentation disorders	Pr : Skin and subcutaneous tissue disorders			

Category: DERMA											
	<p>Adverse Event Pruritus</p> <p>Grade 1: mild or localized, relieved spontaneously or by local measures Grade 2: intense or widespread, relieved spontaneously or by systemic measures Grade 3: intense or widespread and poorly controlled despite treatment N/A Grade 4: Grade 5: death related to toxicity</p>	Pruritus	1, 2 or 3	LLT : Pruritis	PT : Pruritus NOS	HLT : Pruritus NEC	HLGT : Epidermal and dermal conditions	Pr : Skin and subcutaneous tissue disorders	SSC : ANAPHYLAXIS		
	Purpura is graded in the HEMORRHAGE category										
	<p>Adverse Event Radiation dermatitis</p> <p>Grade 1: faint erythema or dry desquamation Grade 2: moderate to brisk erythema or a patchy moist desquamation, mostly confined to skin folds and creases; moderate edema Grade 3: confluent moist desquamation >= 1.5 cm diameter and not confined to skin folds; pitting edema Grade 4: skin necrosis or ulceration of full thickness dermis; may include bleeding not induced by minor trauma or abrasion Grade 5: death related to toxicity</p> <p>Note: Pain associated with radiation dermatitis is graded separately in the PAIN category as Pain due to radiation.</p>	Radiation dermatitis		LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
	<p>Adverse Event Radiation recall reaction (reaction following chemotherapy in the absence of additional radiation therapy that occurs in a previous radiation port)</p> <p>Grade 1: faint erythema or dry desquamation Grade 2: moderate to brisk erythema or a patchy moist desquamation, mostly confined to skin folds and creases; moderate edema Grade 3: confluent moist desquamation >= 1.5 cm diameter and not confined to skin folds; pitting edema Grade 4: skin necrosis or ulceration of full thickness dermis; may include bleeding not induced by minor trauma or abrasion Grade 5: death related to toxicity</p>	Radiation recall reaction		LLT :	PT :	HLT :	HLGT :	Pr :			
	Rash/desquamation		LLT :	PT :	HLT :	HLGT :	Pr :				

<p>Adverse Event Rash/desquamation</p> <p>Grade 1: macular or papular eruption or erythema without associated symptoms Grade 2: macular or papular eruption or erythema with pruritus or other associated symptoms covering <50% of body surface or localized desquamation or other lesions covering <50% of body surface area Grade 3: symptomatic generalized erythroderma or macular, papular or vesicular eruption or desquamation covering >=50% of body surface area Grade 4: generalized exfoliative dermatitis or ulcerative dermatitis Grade 5: death related to toxicity</p> <p>Note: Stevens-Johnson syndrome is graded separately as Erythema multiforme in the DERMATOLOGY/SKIN category. Also consider Allergic reaction/hypersensitivity.</p>	Rash macular		LLT :	PT :	HLT :	HLGT :	Pr :			
	1, 2 or 3	LLT :								
	1, 2 or 3	LLT :		PT :	HLT :	HLGT : Epidermal and dermal conditions	Pr : Skin and subcutaneous tissue disorders			
	Eruption		LLT :	PT :	HLT :	HLGT :	Pr :			
	Rash erythematous	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
		3	LLT :	PT :	HLT :	HLGT :	Pr :			
		3	LLT :	PT :	HLT :	HLGT :	Pr :			
	Dermatitis exfoliative	4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Rash/dermatitis associated with high-dose chemotherapy or BMT studies.</p> <p>Grade 1: faint erythema or dry desquamation Grade 2: moderate to brisk erythema or a patchy moist desquamation, mostly confined to skin folds and creases; moderate edema Grade 3: confluent moist desquamation >1.5 cm diameter and not confined to skin folds; pitting edema Grade 4: skin necrosis or ulceration of full thickness dermis; may include spontaneous bleeding not induced by minor trauma or abrasion N/A Grade 5:</p>			LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Rash/desquamation associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol.</p> <p>Grade 1: macular or papular eruption or erythema covering <25% of body surface area without associated symptoms Grade 2: macular or papular eruption or erythema with pruritus or other</p>	Rash/desquamation		LLT :	PT :	HLT :	HLGT :	Pr :			
	Rash macular		LLT :	PT :	HLT :	HLGT :	Pr :			
	1, 2 or 3	LLT :		PT :	HLT :	HLGT :	Pr :			

<p>Grade 2: macular or papular eruption or erythema with pruritus or other associated symptoms covering ≥ 25 - $<50\%$ of body surface or localized desquamation or other lesions covering ≥ 25 - $<50\%$ of body surface area</p> <p>Grade 3: symptomatic generalized erythroderma or symptomatic macular, papular or vesicular eruption, with bullous formation, or desquamation covering $\geq 50\%$ of body surface area</p> <p>Grade 4: generalized exfoliative dermatitis or ulcerative dermatitis or bullous formation</p> <p>Grade 5: death related to toxicity</p> <p>Note: Stevens-Johnson syndrome is graded separately as Erythema multiforme in the DERMATOLOGY/SKIN category. Also consider Allergic reaction/hypersensitivity.</p>	Papular skin eruption		LLT :	PL :	HLT :	HLGT :	Pr :			
	Eruption		LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
	Erythroderma		LLT :	PT :	HLT :	HLGT :	Pr :			
		3	LLT :	PT :	HLT :	HLGT :	Pr :			
	Dermatitis exfoliative	4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Urticaria (hives, welts, wheals)</p> <p>Grade 1: requiring no medication</p> <p>Grade 2: requiring PO or topical treatment or IV medication or steroids for <24 hours</p> <p>Grade 3: requiring IV medication or steroids for ≥ 24 hours</p> <p>N/A Grade 4:</p> <p>Grade 5: death related to toxicity</p>	Urticaria (hives, welts, wheals)	1, 2 or 3	LLT :			HLGT :	Pr :	SSC: ANAPHYLAXIS		
		1, 2 or 3	LLT :	PT :	HLT :	HLGT : Allergic conditions	Sec ₁ : Immune system disorders			
		1, 2 or 3	LLT :							
	Wheals	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
						HLGT :	Sec ₁ :			
<p>Adverse Event Wound-infectious</p> <p>Grade 1: cellulitis</p> <p>Grade 2: superficial infection</p> <p>Grade 3: infection requiring IV antibiotics</p> <p>Grade 4: necrotizing fasciitis</p> <p>Grade 5: death related to toxicity</p>	Wound-infectious	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
						HLGT :	Sec ₁ :			
	Cellulitis	1	LLT :	PT :	HLT :	HLGT :	Pr :			
						HLGT :	Sec ₁ :			
		4	LLT :	PT :	HLT : Muscle and soft tissue infections	HLGT :	Pr :			
				HLT :	HLGT : Musculoskeletal and connective tissue disorders NEC	Sec ₁ :				

<p>Adverse Event Wound-non-infectious</p> <p>Grade 1: incisional separation Grade 2: incisional hernia Grade 3: fascial disruption without evisceration Grade 4: fascial disruption with evisceration Grade 5: death related to toxicity</p>	Wound-non-infectious		<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p>Adverse Event Dermatology/Skin-Other (Specify,_____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
ENDOCRINE										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Cushingoid appearance (e.g., moon face, buffalo hump, centripetal obesity, cutaneous striae)</p> <p>N/A Grade 1: Grade 2: present N/A Grade 3: N/A Grade 4: N/A Grade 5:</p> <p>Note: Also consider Hyperglycemia, Hypokalemia</p>	Cushingoid appearance (e.g., moon face, buffalo hump, centripetal obesity, cutaneous striae)	2	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
		2	<u>LLT</u> :		<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₁</u> :			
		2	<u>LLT</u> :		<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
		2	<u>LLT</u> :		<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₁</u> :			
<p>Adverse Event Feminization of male</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present N/A Grade 4: Grade 5: death related to toxicity</p>	Feminization of male	3	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
					<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₁</u> :			
<p>Adverse Event Gynecomastia</p>					<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			

Category: ENDOCRINE

<p>Grade 1: mild Grade 2: pronounced or painful Grade 3: pronounced or painful and requiring surgery N/A Grade 4: Grade 5: death related to toxicity</p>	Gynecomastia	1, 2 or 3	LLT : Gynecomastia	PT : Gynaecomastia	HLT :	HLGT :	Sec₁ :				
	<p>Adverse Event Hot flashes/flushes Grade 1: mild or no more than 1 per day Grade 2: moderate and greater than 1 per day N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Hot flashes		LLT :	PT :	HLT :	HLGT :	Pr :			
		Hot flashes		LLT :							
			1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
	<p>Adverse Event Hypothyroidism Grade 1: asymptomatic, TSH elevated, no therapy given Grade 2: symptomatic or thyroid replacement treatment given Grade 3: patient hospitalized for manifestations of hypothyroidism Grade 4: myxedema coma Grade 5: death related to toxicity</p>	Hypothyroidism		LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			
			1	LLT :	PT :	HLT :	HLGT :	Pr :			
		Myxedema	4	LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			
						HLT :	HLGT :	Pr :			
	<p>Adverse Event Masculinization of female N/A Grade 1: N/A Grade 2: Grade 3: present N/A Grade 4: Grade 5: death related to toxicity</p>	<p>3</p> <p>Masculinization of female</p>		LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			
	<p>Adverse Event SIADH (syndrome of inappropriate antidiuretic hormone) N/A Grade 1: N/A Grade 2: Grade 3: present N/A Grade 4: Grade 5: death related to toxicity</p>	<p>SIADH (syndrome of inappropriate antidiuretic hormone)</p>		LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			

<p>Adverse Event Endocrine-Other (Specify, _____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
GASTROINTESTINAL										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
Amylase is graded in the METABOLIC/LABORATORY category										
<p>Adverse Event Anorexia</p> <p>Grade 1: loss of appetite Grade 2: oral intake significantly decreased Grade 3: requiring IV fluids Grade 4: requiring feeding tube or parenteral nutrition Grade 5: death related to toxicity</p>	Anorexia	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
		1	<u>LLT</u> :							
<p>Adverse Event Ascites (non-malignant)</p> <p>Grade 1: asymptomatic Grade 2: symptomatic, requiring diuretics Grade 3: symptomatic, requiring therapeutic paracentesis Grade 4: life-threatening physiologic consequences Grade 5: death related to toxicity</p>	Ascites (non-malignant)	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
					<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₁</u> :			
					<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₂</u> :			

<p>Adverse Event Colitis</p> <p><u>N/A Grade 1:</u> <u>Grade 2:</u> abdominal pain with mucus and/or blood in stool <u>Grade 3:</u> abdominal pain, fever, change in bowel habits with ileus or peritoneal signs, and radiographic or biopsy documentation <u>Grade 4:</u> perforation or requiring surgery or toxic megacolon <u>Grade 5:</u> death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia, Melena/GI bleeding, Rectal bleeding/hematochezia, Hypotension.</p>	Colitis		<u>LLT :</u>	<u>PT :</u>	<u>HLT :</u>	<u>HLGT :</u>	<u>Pr :</u>			
<p>Adverse Event Constipation</p> <p><u>Grade 1:</u> requiring stool softener or dietary modification <u>Grade 2:</u> requiring laxatives <u>Grade 3:</u> obstipation requiring manual evacuation or enema <u>Grade 4:</u> obstruction or toxic megacolon <u>Grade 5:</u> death related to toxicity</p>	Constipation	1, 2, 3 or 4	<u>LLT :</u>	<u>PT :</u>	<u>HLT :</u>	<u>HLGT :</u>	<u>Pr :</u>			
<p>Adverse Event Dehydration</p> <p><u>Grade 1:</u> dry mucous membranes and/or diminished skin turgor <u>Grade 2:</u> requiring IV fluid replacement (brief) <u>Grade 3:</u> requiring IV fluid replacement (sustained) <u>Grade 4:</u> physiologic consequences requiring intensive care; or hemodynamic collapse <u>Grade 5:</u> death related to toxicity</p> <p>Note: Also consider Diarrhea, Vomiting, Stomatitis/pharyngitis (oral/pharyngeal mucositis), Hypotension.</p>	Dehydration	1, 2, 3 or 4	<u>LLT :</u>	<u>PT :</u>	<u>HLT :</u>	<u>HLGT :</u>	<u>Pr :</u>			
		1	<u>LLT :</u>	<u>PT :</u>	<u>HLT :</u>	<u>HLGT :</u>	<u>Pr :</u>			

<p>Adverse Event Diarrhea patients without colostomy</p> <p>Grade 1: increase of <4 stools/day over pre-treatment Grade 2: increase of 4-6 stools/day, or nocturnal stools Grade 3: increase of >= 7 stools/day or incontinence; or need for parenteral support for dehydration Grade 4: physiologic consequences requiring intensive care; or hemodynamic collapse Grade 5: death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia, Pain, Dehydration, Hypotension.</p>	<p>Diarrhea patients without colostomy</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Diarrhea patients with a colostomy</p> <p>Grade 1: mild increase in loose, watery colostomy output compared with pretreatment Grade 2: moderate increase in loose, watery colostomy output compared with pretreatment, but not interfering with normal activity Grade 3: severe increase in loose, watery colostomy output compared with pretreatment, interfering with normal activity Grade 4: physiologic consequences, requiring intensive care; or hemodynamic collapse Grade 5: death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia, Pain, Dehydration, Hypotension.</p>	<p>Diarrhea patients with a colostomy</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Diarrhea associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol.</p> <p>Grade 1: > 500 - <= 1000 mL of diarrhea/day Grade 2: > 1000 - <=1500 mL of diarrhea/day Grade 3: > 1500 mL of diarrhea/day Grade 4: severe abdominal pain with or without ileus Grade 5: death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia, Pain, Dehydration, Hypotension.</p>	<p>Diarrhea associated with graft versus host disease (GVHD)</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
			<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Sec₁ :</p>	<p>SSC₁ : SECONDARY IMMUNOCOM PROMISED STATE</p>		

<p>Adverse Event Diarrhea for pediatric BMT studies, if specified in the protocol.</p> <p>Grade 1: >5 - <=10 mL/kg of diarrhea/day Grade 2: >10 - <= 15 mL/kg of diarrhea/day Grade 3: >15 mL/kg of diarrhea/day N/A Grade 4: Grade 5: death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia, Pain, Dehydration, Hypotension.</p>	Diarrhea	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Duodenal ulcer (requires radiographic or endoscopic documentation)</p> <p>N/A Grade 1: Grade 2: requiring medical management or non-surgical treatment Grade 3: uncontrolled by outpatient medical management; requiring hospitalization Grade 4: perforation or bleeding, requiring emergency surgery Grade 5: death related to toxicity</p>	Duodenal ulcer	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
<p>Adverse Event Dyspepsia/heartburn</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe N/A Grade 4: N/A Grade 5:</p>	Dyspepsia/heartburn	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Dysphagia, esophagitis, odynophagia (painful swallowing)</p> <p>Grade 1: mild dysphagia, but can eat regular diet Grade 2: dysphagia, requiring predominantly pureed, soft, or liquid diet Grade 3: dysphagia, requiring IV hydration Grade 4: complete obstruction (cannot swallow saliva) requiring enteral or parenteral nutritional support, or perforation Grade 5: death related to toxicity</p> <p>Note: If the adverse event is radiation-related, grade either under Dysphagia-esophageal related to radiation or Dysphagia-pharyngeal related to radiation.</p>	Dysphagia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Dysphagia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Odynophagia		LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			

<p>Adverse Event Dysphagia-esophageal related to radiation</p> <p>Grade 1: mild dysphagia, but can eat regular diet Grade 2: dysphagia, requiring predominantly pureed, soft, or liquid diet. Grade 3: dysphagia, requiring feeding tube, IV hydration, or hyperalimentation Grade 4: complete obstruction (cannot swallow saliva); ulceration with bleeding not induced by minor trauma or abrasion or perforation Grade 5: death related to toxicity</p> <p>Note: Fistula is graded separately as Fistula-esophageal. Also consider Pain due to radiation, Mucositis due to radiation.</p>	<p>Dysphagia-esophageal related to radiation</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Dysphagia-pharyngeal related to radiation</p> <p>Grade 1: mild dysphagia, but can eat regular diet Grade 2: dysphagia, requiring predominantly pureed, soft, or liquid diet. Grade 3: dysphagia, requiring feeding tube, IV hydration, or hyperalimentation Grade 4: complete obstruction (cannot swallow saliva); ulceration with bleeding not induced by minor trauma or abrasion or perforation Grade 5: death related to toxicity</p> <p>Note: Fistula is graded separately as Fistula-esophageal. Also consider Pain due to radiation, Mucositis due to radiation.</p>	<p>Dysphagia-pharyngeal related to radiation</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Fistula-esophageal</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present Grade 4: requiring surgery Grade 5: death related to toxicity</p>	<p>Fistula-esophageal</p>	<p>3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Fistula-intestinal</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present Grade 4: requiring surgery Grade 5: death related to toxicity</p>	<p>Fistula-intestinal</p>	<p>3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			

Category: GASTROINTESTINAL

<p>Adverse Event Fistula-Pharyngeal</p> <p><u>N/A Grade 1:</u> <u>N/A Grade 2:</u> <u>Grade 3:</u> present <u>Grade 4:</u> requiring surgery <u>Grade 5:</u> death related to toxicity</p>	Fistula-Pharyngeal		<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p>Adverse Event Fistula-rectal/anal</p> <p><u>N/A Grade 1:</u> <u>N/A Grade 2:</u> <u>Grade 3:</u> present <u>Grade 4:</u> requiring surgery <u>Grade 5:</u> death related to toxicity</p>		3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
		3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p>Adverse Event Flatulence</p> <p><u>Grade 1:</u> mild <u>Grade 2:</u> moderate <u>N/A Grade 3:</u> <u>N/A Grade 4:</u> <u>N/A Grade 5:</u></p>	Flatulence	1 or 2	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p>Adverse Event Gastric ulcer (requires radiographic or endoscopic documentation)</p> <p><u>N/A Grade 1:</u> <u>Grade 2:</u> requiring medical management or non-surgical treatment <u>Grade 3:</u> bleeding without perforation, uncontrolled by outpatient medical management; requiring hospitalization or surgery <u>Grade 4:</u> perforation or bleeding, requiring emergency surgery <u>Grade 5:</u> death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia</p> <p>Attribution Code Unrelated Unlikely Possible Probable Definite</p>	Gastric ulcer	2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :	<u>SSC</u> : UPPER GI BLEEDING/PERFORATION		

<p>Adverse Event Gastritis</p> <p>N/A Grade 1: Grade 2: requiring medical management or non-surgical treatment Grade 3: uncontrolled by out-patient medical management; requiring hospitalization or surgery Grade 4: life-threatening bleeding, requiring emergency surgery Grade 5: death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia</p>	Gastritis	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Hematemesis is graded in the HEMORRHAGE category</p>										
<p>Hematochezia is graded in the HEMORRHAGE category as Rectal bleeding/hematochezia</p>										
<p>Adverse Event Ileus (or neuroconstipation)</p> <p>N/A Grade 1: Grade 2: intermittent, not requiring intervention Grade 3: requiring non-surgical intervention Grade 4: requiring surgery Grade 5: death related to toxicity</p>	Ileus (or neuroconstipation)	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event mouth dryness</p> <p>Grade 1: mild Grade 2: moderate N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Mouth dryness		LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Note: Mucositis not due to radiation is graded in the GASTROINTESTINAL category for specific sites: Colitis, Esophagitis, Gastritis, Stomatitis/pharyngitis (oral/pharyngeal mucositis), and Typhlitis; or the RENAL/GENITOURINARY category for Vaginitis</p>										

<p>Adverse Event Mucositis due to radiation</p> <p>Grade 1: erythema of the mucosa Grade 2: patchy pseudomembranous reaction (patches generally <= 1.5 cm in diameter and non-contiguous) Grade 3: confluent pseudomembranous reaction (contiguous patches generally > 1.5 cm in diameter) Grade 4: necrosis or deep ulceration; may include bleeding not induced by minor trauma or abrasion Grade 5: death related to toxicity</p> <p>Note: Grade radiation mucositis of the larynx here. Dysphagia related to radiation is also graded as either Dysphagia-esophageal related to radiation or Dysphagia-pharyngeal related to radiation, depending on the site of treatment. Also consider Pain due to radiation.</p>	<p>Mucositis due to radiation</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Nausea</p> <p>Grade 1: able to eat Grade 2: oral intake significantly decreased Grade 3: no significant intake, requiring IV fluids N/A Grade 4: Grade 5: death related to toxicity</p> <p>Attribution Code Unrelated Unlikely Possible Probable Definite</p>	<p>Nausea</p>	<p>1, 2 or 3</p>	<p>LLT : Nausea</p>	<p>PT : Nausea</p>	<p>HLT : Nausea and vomiting symptoms</p>	<p>HLGT : Gastrointestinal signs and symptoms</p>	<p>Pr : Gastrointestinal disorders</p>			
<p>Adverse Event Pancreatitis</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: abdominal pain with pancreatic enzyme elevation Grade 4: complicated by shock (acute circulatory failure) Grade 5: death related to toxicity</p> <p>Note: Amylase is graded in the METABOLIC/LABORATORY category. Also consider Hypotension.</p>	<p>Pancreatitis</p>	<p>3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Pharyngitis is graded in the GASTROINTESTINAL category as Stomatitis/pharyngitis (oral/pharyngeal mucositis)</p>										

<p>Adverse Event Proctitis</p> <p>Grade 1: increased stool frequency, occasional blood-streaked stools or rectal discomfort (including hemorrhoids) not requiring medication Grade 2: increased stool frequency, bleeding, mucus discharge, or rectal discomfort requiring medication; anal fissure Grade 3: increased stool frequency/diarrhea requiring parenteral support; rectal bleeding requiring transfusion; or persistent mucus discharge, necessitating pads Grade 4: perforation, bleeding or necrosis or other life-threatening complication requiring surgical intervention (e.g., colostomy) Grade 5: death related to toxicity</p> <p>Note: Proctitis occurring more than 90 days after the start of radiation therapy is graded in the RTOG/EORTC Late Radiation Morbidity Scoring Scheme (See Appendix IV). Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia, Pain due to radiation. Fistula is graded separately as Fistula-rectal/anal.</p>	Proctitis		LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Salivary gland changes</p> <p>Grade 1: slightly thickened saliva; may have slightly altered taste (e.g., metallic); additional fluids may be required Grade 2: thick, ropy, sticky saliva; markedly altered taste; alteration in diet required N/A Grade 3: Grade 4: acute salivary gland necrosis Grade 5: death related to toxicity</p>	Salivary gland changes	1, 2 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Sense of smell</p> <p>Grade 1: slightly altered Grade 2: markedly altered N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Sense of smell	1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Stomatitis/pharyngitis (oral/pharyngeal mucositis)</p> <p>Grade 1: painless ulcers, erythema, or mild soreness in the absence of lesions Grade 2: painful erythema, edema, or ulcers, but can eat or swallow Grade 3: painful erythema, edema, or ulcers requiring IV hydration Grade 4: severe ulceration or requires parenteral or enteral nutritional support or prophylactic intubation Grade 5: death related to toxicity</p>	Stomatitis/pharyngitis (oral/pharyngeal mucositis)	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Pharyngitis	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			

<p>Note: Radiation-related mucositis is graded as Mucositis due to radiation.</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Stomatitis/pharyngitis (oral/pharyngeal mucositis) for BMT studies, if specified in the protocol.</p> <p>Grade 1: painless ulcers, erythema, or mild soreness in the absence of lesions Grade 2: painful erythema, edema or ulcers but can swallow Grade 3: painful erythema, edema, or ulcers preventing swallowing or requiring hydration or parenteral (or enteral) nutritional support Grade 4: severe ulceration requiring prophylactic intubation or resulting in documented aspiration pneumonia Grade 5: death related to toxicity</p> <p>Note: Radiation-related mucositis is graded as Mucositis due to radiation.</p>	Stomatitis/pharyngitis (oral/pharyngeal mucositis)	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Taste disturbance (dysgeusia)</p> <p>Grade 1: slightly altered Grade 2: markedly altered N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Taste disturbance	1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
	Dysgeusia	1 or 2	LLT :	PT :	HLT :	HLGT :	Sec :			
<p>Adverse Event Typhilitis (inflammation of cecum)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: abdominal pain, diarrhea, fever, and radiographic or biopsy documentation Grade 4: perforation, bleeding or necrosis or other life-threatening complication requiring surgical intervention (e.g., colostomy) Grade 5: death related to toxicity</p> <p>Note: Also consider Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia, Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia, Hypotension, Febrile neutropenia.</p>		3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Inflammation of cecum		LLT :							

<p><u>Adverse Event</u> Vomiting</p> <p><u>Grade 1</u>: 1 episode in 24 hours over pretreatment <u>Grade 2</u>: 2-5 episodes in 24 hours over pretreatment <u>Grade 3</u>: >= 6 episodes in 24 hours over pretreatment; or need for IV fluids <u>Grade 4</u>: requiring parenteral nutrition; or physiologic consequences requiring intensive care; hemodynamic collapse <u>Grade 5</u>: death related to toxicity</p> <p>Note: Also consider Dehydration.</p>	Vomiting	1, 2, 3 or 4	<u>LLT</u> : Vomiting	<u>PT</u> : Vomiting NOS	<u>HLT</u> : Nausea and vomiting symptoms	<u>HLGT</u> : Gastrointestinal signs and symptoms	<u>Pr</u> : Gastrointestinal disorders			
Weight gain is graded in the CONSTITUTIONAL SYMPTOMS category										
Weight loss is graded in the CONSTITUTIONAL SYMPTOMS category										
<p><u>Adverse Event</u> Gastrointestinal-Other (Specify, _____)</p> <p><u>Grade 1</u>: mild <u>Grade 2</u>: moderate <u>Grade 3</u>: severe <u>Grade 4</u>: life-threatening or disabling <u>Grade 5</u>: death related to toxicity</p>										
HEMORRHAGE										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Note: Transfusion in this section refers to pRBC infusion. For any bleeding with grade 3 or 4 platelets (< 50,000), always grade Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia. Also consider Platelets, Transfusion: pRBCs, and Transfusion: Platelets in addition to grading severity by grading the site or type of bleeding. If the site or type of Hemorrhage/bleeding is listed, also use the grading that incorporates the site of bleeding: CNS Hemorrhage/bleeding, Hematuria, Hematemesis, Hemoptysis, Hemorrhage/bleeding with surgery, Melena/GI bleeding, Petechiae/purpura (Hemorrhage/bleeding into skin), Rectal bleeding/hematochezia, Vaginal bleeding. If the platelet count is >= 50,000 and the site or type of bleeding is listed, grade the specific site. If the site or type is not listed and the platelet count is >= 50,000, grade Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia and specify the site or type in the OTHER category.</p>										

<p>Adverse Event Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia</p> <p>Grade 1: mild without transfusion N/A Grade 2: Grade 3: requiring transfusion Grade 4: catastrophic bleeding, requiring major non-elective intervention Grade 5: death related to toxicity</p> <p>Note: If the site is not listed, grade as Hemorrhage-Other (Specify site, _____). This adverse event must be graded for any bleeding with grade 3 or 4 thrombocytopenia. Also consider Platelets, Hemoglobin, Transfusion: platelets, Transfusion: pRBCs, site or type of bleeding.</p>	<p>Hemorrhage/bleeding with grade 3 or 4 thrombocytopenia</p>	<p>1, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>	<p>SSC₁ :</p>		
<p>LLT :</p> <p>PT :</p> <p>HLT :</p> <p>HLGT :</p> <p>Pr :</p> <p>SSC₁ :</p>										
<p>Adverse Event Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia</p> <p>Grade 1: mild without transfusion N/A Grade 2: Grade 3: requiring transfusion Grade 4: catastrophic bleeding requiring major non-elective intervention Grade 5: death related to toxicity</p> <p>Note: Bleeding in the absence of grade 3 or 4 thrombocytopenia is graded here only if the specific site or type of bleeding is not listed elsewhere in the HEMORRHAGE category. Also grade as Other in the HEMORRHAGE category. Also consider Platelets, Hemoglobin, Transfusion: platelets, Transfusion: pRBCs, Hemorrhage - Other (Specify site, _____).</p>	<p>Hemorrhage/bleeding without grade 3 or 4 thrombocytopenia</p>	<p>1, 3 or 4</p>	<p>If you need to set a complete mapping version (CTC versions 2.0 and/or 3.0 - MedDRA), please contact Pharmadhoc 112, rue Olivier de Serres 75015 Paris Tel: +33 (0)8 71 76 88 17 Tel: +33 (0)6 73 51 37 96 or send an email to info@pharmadhoc.com</p>					<p>SSC₁ :</p>		
<p>Adverse Event CNS hemorrhage/bleeding</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: bleeding noted on CT or other scan with no clinical consequences Grade 4: hemorrhagic stroke or hemorrhagic vascular event (CVA) with neurologic signs and symptoms Grade 5: death related to toxicity</p>	<p>CNS hemorrhage/bleeding</p>	<p>3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>	<p>SSC₁ :</p>		
<p>LLT :</p> <p>PT :</p> <p>HLT :</p> <p>HLGT :</p> <p>Pr :</p> <p>SSC₁ :</p>										
<p>Adverse Event Epistaxis</p> <p>Grade 1: mild without transfusion N/A Grade 2:</p>	<p>Epistaxis</p>		<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>	<p>SSC₁ :</p>		

Category: HEMORRHAGE

<p><u>Grade 3</u>: requiring transfusion <u>Grade 4</u>: catastrophic bleeding, requiring major non-elective intervention <u>Grade 5</u>: death related to toxicity</p>	Epistaxis		<u>LLT</u>	<u>PT</u>	<u>HLT</u>	<u>HLGT</u>	<u>Sec₁</u>	<u>SSC₁</u>		
<p><u>Adverse Event</u> Hematemesis <u>Grade 1</u>: mild without transfusion <u>N/A Grade 2</u>: <u>Grade 3</u>: requiring transfusion <u>Grade 4</u>: catastrophic bleeding, requiring major non-elective intervention <u>Grade 5</u>: death related to toxicity</p>	Hematemesis	1, 3 or 4	<u>LLT</u>	<u>PT</u>	<u>HLT</u>	<u>HLGT</u>	<u>Pr</u>	<u>SSC₁</u>	<u>SSC₂</u>	
					<u>HLT</u>	<u>HLGT</u>	<u>Sec₁</u>			
<p><u>Adverse Event</u> Hematuria (in the absence of vaginal bleeding) <u>Grade 1</u>: microscopic only <u>Grade 2</u>: intermittent gross bleeding, no clots <u>Grade 3</u>: persistent gross bleeding or clots; may require catheterization or instrumentation, or transfusion <u>Grade 4</u>: open surgery or necrosis or deep bladder ulceration <u>Grade 5</u>: death related to toxicity</p>	Hematuria (in the absence of vaginal bleeding)	1, 2, 3 or 4	<u>LLT</u>	<u>PT</u>	<u>HLT</u>	<u>HLGT</u>	<u>Pr</u>	<u>SSC₁</u>		
					<u>HLT</u>	<u>HLGT</u>	<u>Sec₁</u>			
<p><u>Adverse Event</u> Hemoptysis <u>Grade 1</u>: mild without transfusion <u>N/A Grade 2</u>: <u>Grade 3</u>: requiring transfusion <u>Grade 4</u>: catastrophic bleeding, requiring major non-elective intervention <u>Grade 5</u>: death related to toxicity</p>		1, 3 or 4	<u>LLT</u>	<u>PT</u>	<u>HLT</u>	<u>HLGT</u>	<u>Pr</u>	<u>SSC₁</u>		
					<u>HLT</u>	<u>HLGT</u>	<u>Sec₁</u>			
					<u>HLT</u> : Haemorrhages NEC	<u>HLGT</u> : Vascular haemorrhagic disorders	<u>Sec₂</u> : Vascular disorders			
<p><u>Adverse Event</u> Hemorrhage/bleeding associated with surgery <u>Grade 1</u>: mild without transfusion <u>N/A Grade 2</u>: <u>Grade 3</u>: requiring transfusion <u>Grade 4</u>: catastrophic bleeding, requiring major non-elective intervention <u>Grade 5</u>: death related to toxicity</p> <p>Note: Expected blood loss at the time of surgery is not graded as an adverse event.</p>	Hemorrhage/bleeding associated with surgery	1, 3 or 4	<u>LLT</u>	<u>PT</u>	<u>HLT</u>	<u>HLGT</u>	<u>Pr</u>			

<p>Adverse Event Melena/GI bleeding</p> <p>Grade 1: mild without transfusion N/A Grade 2: Grade 3: requiring transfusion Grade 4: catastrophic bleeding, requiring major non-elective intervention Grade 5: death related to toxicity</p>	Melena/GI bleeding	1, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : HAEMORRHA GE	SSC₂ : UPPER GI BLEEDING/P ERFORATIO N	
		1, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : Gastrointestinal disorders	SSC₁ : HAEMORRHA GE		
<p>Adverse Event Petechiae/purpura (hemorrhage/bleeding into skin or mucosa)</p> <p>Grade 1: rare petechiae of skin Grade 2: petechiae or purpura in dependent areas of skin Grade 3: generalized petechiae or purpura of skin or petechiae of any mucosal site N/A Grade 4: Grade 5: death related to toxicity</p>	Petechiae/purpura (hemorrhage/bleeding into skin or mucosa)		LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
	Purpura	1, 2 or 3	LLT : Purpura	PT : Purpura NOS	HLT : Purpura and related conditions HLT : Purpuras (excl thrombocytopenic) HLT : Bruising, ecchymosis and purpura	HLGT : Skin vascular abnormalities HLGT : Bleeding tendencies and purpuras (excl thrombocytopenic) HLGT : Vascular haemorrhagic disorders	Pr : Skin and subcutaneous tissue disorders Sec₁ : Blood and lymphatic system disorders Sec₂ : Vascular disorders	SSC₁ : HAEMORRHA GE		
<p>Adverse Event Rectal bleeding/hematochezia</p> <p>Grade 1: mild without transfusion or medication Grade 2: persistent, requiring medication (e.g., steroid suppositories) and/or break from radiation treatment Grade 3: requiring transfusion Grade 4: catastrophic bleeding, requiring major non-elective intervention Grade 5: death related to toxicity</p>	Rectal bleeding/hematochezia		LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :		
	Hematochezia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	Sec₁ :		
<p>Adverse Event Vaginal bleeding</p> <p>Grade 1: spotting, requiring <2 pads per day Grade 2: requiring >= 2 pads per day, but not requiring transfusion Grade 3: requiring transfusion Grade 4: catastrophic bleeding, requiring major non-elective intervention Grade 5: death related to toxicity</p>	Vaginal bleeding	1, 2, 3 or 4	LLT : Vaginal bleeding	PT : Vaginal haemorrhage	HLT :	HLGT :	Pr :	SSC₁ : HAEMORRHA GE		
					HLT : Reproductive system haemorrhages	HLGT : Vascular haemorrhagic disorders	Sec₁ : Vascular disorders			

<p>Adverse Event Hemorrhage-Other (Specify, _____)</p> <p>Grade 1: mild without transfusion N/A Grade 2: Grade 3: requiring transfusion Grade 4: catastrophic bleeding, requiring major non-elective intervention Grade 5: death related to toxicity</p>										
HEPATIC										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Alkaline phosphatase</p> <p>Grade 1: > ULN - 2.5 x ULN Grade 2: > 2.5 - 5.0 x ULN Grade 3: > 5.0 - 20.0 x ULN Grade 4: > 20.0 x ULN Grade 5: death related to toxicity</p>	Alkaline phosphatase	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Bilirubin</p> <p>Grade 1: > ULN - 1.5 x ULN Grade 2: > 1.5 - 3.0 x ULN Grade 3: > 3.0 - 10.0 x ULN Grade 4: > 10.0 x ULN Grade 5: death related to toxicity</p>	Bilirubin	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Bilirubin associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol.</p> <p>Grade 1: >= 2 - <3 mg/100 mL Grade 2: >= 3 - <6 mg/100 mL Grade 3: >= 6 - <15 mg/100 mL Grade 4: >= 15 mg/100 mL Grade 5: death related to toxicity</p>	Bilirubin associated with graft versus host disease (GVHD)	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT :	Pr :	SSC_n :		
					HLT :	HLGT :	Sec_n :			

Category: HEPATIC

<p>Adverse Event GGT (Gamma-Glutamyl transpeptidase)</p> <p>Grade 1: > ULN - 2.5 x ULN Grade 2: > 2.5 - 5.0 x ULN Grade 3: > 5.0 - 20.0 x ULN Grade 4: > 20.0 x ULN Grade 5: death related to toxicity</p>	GGT	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hepatic enlargement</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present N/A Grade 4: Grade 5: death related to toxicity</p> <p>Note: Grade Hepatic enlargement only for treatment related adverse event including Veno-Occlusive Disease.</p>	Hepatic enlargement	3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hypoalbuminemia</p> <p>Grade 1: < LLN - 3 g/dL Grade 2: >= 2 - <3 g/dL Grade 3: <2 g/dL N/A Grade 4: Grade 5: death related to toxicity</p>	Hypoalbuminemia	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Liver dysfunction/failure (clinical)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: asterixis Grade 4: encephalopathy or coma Grade 5: death related to toxicity</p>	Liver dysfunction/failure	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		3	LLT :	PT :	HLT :	HLGT : Movement disorders (incl Parkinsonism)	Pr :			
	Hepatic encephalopathy	4	LLT :	PT :	HLT :	HLGT : Hepatic and hepatobiliary disorders	Sec₁ : Hepatobiliary disorders			
		4	LLT :	PT :	HLT :	HLGT :	Pr :			

			LLT	PL	HLT :	HLGT :	Sec ₁ :			
<p>Adverse Event Portal vein flow</p> <p>N/A Grade 1: Grade 2: decreased portal vein flow Grade 3: reversal/retrograde portal vein flow N/A Grade 4: Grade 5: death related to toxicity</p>	Portal vein flow		LLT :	PT :	HLT :	HLGT :	Pr :			
				LLT :		HLT :	HLGT :	Sec ₁ :		
<p>Adverse Event SGOT (AST) (serum glutamic oxaloacetic transaminase)</p> <p>Grade 1: > ULN - 2.5 x ULN Grade 2: > 2.5 - 5.0 x ULN Grade 3: > 5.0 - 20.0 x ULN Grade 4: > 20.0 x ULN Grade 5: death related to toxicity</p>	SGOT	1, 2, 3 or 4	LLT :							
			1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :		
<p>Adverse Event SGPT (ALT) (serum glutamic pyruvic transaminase)</p> <p>Grade 1: > ULN - 2.5 x ULN Grade 2: > 2.5 - 5.0 x ULN Grade 3: > 5.0 - 20.0 x ULN Grade 4: > 20.0 x ULN Grade 5: death related to toxicity</p>	SGPT	1, 2, 3 or 4	LLT :							
	ALT		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :		
<p>Adverse Event Hepatic-Other (Specify,_____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
INFECTION/FEBRILE NEUTROPENIA										

	Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
ECTION/FEBRILE NEUTROPENIA	<p>Adverse Event Catheter-related infection</p> <p>Grade 1: mild, no active treatment Grade 2: moderate, localized infection, requiring local or oral treatment Grade 3: severe, systemic infection, requiring IV antibiotic or antifungal treatment or hospitalization Grade 4: life-threatening sepsis (e.g., septic shock) Grade 5: death related to toxicity</p>	Catheter-related infection	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr : I			
						HLT :	HLGT :	Sec ₁ : General disorders and administration site conditions			
						HLT :	HLGT :	Sec ₂ : Injury, poisoning and procedural complications			
	<p>Adverse Event Febrile neutropenia (fever of unknown origin without clinically or microbiologically documented infection) (ANC <1.0 x 10e9/L, fever >=38.5 degrees C)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present Grade 4: life-threatening sepsis (e.g., septic shock) Grade 5: death related to toxicity</p> <p>Note: Hypothermia instead of fever may be associated with neutropenia and is graded here. Also consider Neutrophils.</p>		3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT : Febrile disorders	HLGT :	Sec ₁ : General disorders and administration site conditions			
	<p>Adverse Event Infection (documented clinically or microbiologically) with grade 3 or 4 neutropenia (ANC <1.0 x 10e9/L)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present Grade 4: life-threatening sepsis (e.g., septic shock) Grade 5: death related to toxicity</p> <p>Note: Also consider Neutrophils. Hypothermia instead of fever may be associated with neutropenia and is graded here. In the absence of documented infection grade 3 or 4 neutropenia with fever is graded as Febrile neutropenia.</p>	Infection with neutropenia	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
HLT :						HLGT :	Sec ₁ :				

Category: INFI	<p>Adverse Event Infection with unknown ANC</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present Grade 4: life-threatening sepsis (e.g., septic shock) Grade 5: death related to toxicity</p> <p>Note: This adverse event criterion is used in the rare case when ANC is unknown.</p>	Infection	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
	<p>Adverse Event Infection without neutropenia</p> <p>Grade 1: mild, no active treatment Grade 2: moderate localized infection, requiring local or oral treatment Grade 3: severe, systemic infection, requiring IV antibiotic or antifungal treatment, or hospitalization Grade 4: life-threatening sepsis (e.g., septic shock) Grade 5: death related to toxicity</p> <p>Note: Also consider Neutrophils.</p>	Infection without neutropenia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
	<p>Wound-infectious is graded in the DERMATOLOGY/SKIN category</p>											
	<p>Adverse Event Infection/Febrile Neutropenia-Other (Specify,____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>											
LYMPHATICS												
	Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3	
	Adverse Event Lymphatics	Lymphatics	1, 2, 3 or 4	LLT :	PT :	HLT : Lymphangiopathies	HLGT : Lymphatic noninfective disorders	Pr : Vascular disorders				

Category: LYMPHATICS	<p>Grade 1: mild lymphedema Grade 2: moderate lymphedema requiring compression; lymphocyst Grade 3: severe lymphedema limiting function; lymphocyst requiring surgery Grade 4: severe lymphedema limiting function with ulceration Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC : OEDEMA		
	<p>Adverse Event Lymphatics-Other (Specify,____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										

METABOLIC/LABORATORY

Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3	
<p>Adverse Event Acidosis (metabolic or respiratory)</p> <p>Grade 1: pH < normal, but >= 7.3 N/A Grade 2: Grade 3: pH <7.3 Grade 4: pH <7.3 with life-threatening physiologic consequences Grade 5: death related to toxicity</p>	Acidosis	1, 3 or 4	LLT : Acidosis	PT : Acidosis NOS	HLT : Metabolic acidoses (excl diabetic acidoses)	HLGT : Acid-base disorders	Pr : metabolism and nutrition disorders				
			LLT : Acidosis metabolic	PT : Metabolic acidosis NOS	HLT : Metabolic acidoses (excl diabetic acidoses)	HLGT : Acid-base disorders	Pr : metabolism and nutrition disorders				
				LLT :	PT :	HLT : Conditions associated with abnormal gas exchange	HLGT : Respiratory disorders NEC	Pr : Respiratory, thoracic and mediastinal disorders			
						HLT : Respiratory acidoses	HLGT : Acid-base disorders	Sec₁ : Metabolism and nutrition disorders			
<p>Adverse Event Alkalosis (metabolic or respiratory)</p> <p>Grade 1: pH > normal, but <= 7.5 N/A Grade 2: Grade 3: pH > 7.5 Grade 4: pH > 7.5 with life-threatening physiologic consequences Grade 5: death related to toxicity</p>		1, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
		1, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
			LLT :	PT :	HLT :	HLGT :	Pr :				
					HLT :	HLGT :	Sec₁ :				

<p>Adverse Event Amylase</p> <p>Grade 1: >ULN - 1.5 x ULN Grade 2: >1.5 - 2.0 x ULN Grade 3: >2.0 - 5.0 x ULN Grade 4: >5.0 x ULN Grade 5: death related to toxicity</p>	Amylase	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Bicarbonate</p> <p>Grade 1: < LLN - 16 mEq/dL Grade 2: 11 - 15 mEq/dL Grade 3: 8 - 10 mEq/dL Grade 4: <8 mEq/dL Grade 5: death related to toxicity</p>	Bicarbonate	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event CPK (creatine phosphokinase)</p> <p>Grade 1: >ULN - 2.5 x ULN Grade 2: > 2.5 - 5 x ULN Grade 3: > 5 - 10 x ULN Grade 4: > 10 x ULN Grade 5: death related to toxicity</p>	CPK	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
<p>Adverse Event Hypercalcemia</p> <p>Grade 1: >ULN - 11.5 mg/dL; >ULN - 2.9 mmol/L Grade 2: >11.5 - 12.5 mg/dL; >2.9 - 3.1 mmol/L Grade 3: >12.5 - 13.5 mg/dL; >3.1 - 3.4 mmol/L Grade 4: > 13.5 mg/dL; >3.4 mmol/L Grade 5: death related to toxicity</p>	Hypercalcemia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
					HLT :	HLGT :	Sec₁ :			

TORY

<p><u>Adverse Event</u> Hypercholesterolemia</p> <p><u>Grade 1</u>: >ULN - 300 mg/dL ; >ULN - 7.75 mmol/L <u>Grade 2</u>: >300 - 400 mg/dL ; >7.75 - 10.34 mmol/L <u>Grade 3</u>: >400 - 500 mg/dL ; >10.34 - 12.92 mmol/L <u>Grade 4</u>: >500 mg/dL ; >12.92 mmol/L <u>Grade 5</u>: death related to toxicity</p>	Hypercholesterolemia	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p><u>Adverse Event</u> Hyperglycemia</p> <p><u>Grade 1</u>: >ULN - 160 mg/dL ; >ULN - 8.9 mmol/L <u>Grade 2</u>: >160 - 250 mg/dL ; >8.9 - 13.9 mmol/L <u>Grade 3</u>: >250 - 500 mg/dL ; >13.9 - 27.8 mmol/L <u>Grade 4</u>: >500 mg/dL ; >27.8 mmol/L or acidosis <u>Grade 5</u>: death related to toxicity</p>	Hyperglycemia	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> : Hyperglycaemic conditions NEC	<u>HLGT</u> : Glucose metabolism disorders (incl diabetes mellitus)	<u>Pr</u> : metabolism and nutrition disorders <u>Sec</u> ₁ : Endocrine disorders			
<p><u>Adverse Event</u> Hyperkalemia</p> <p><u>Grade 1</u>: >ULN - 5.5 mmol/L <u>Grade 2</u>: >5.5 - 6.0 mmol/L <u>Grade 3</u>: >6.0 - 7.0 mmol/L <u>Grade 4</u>: >7.0 mmol/L <u>Grade 5</u>: death related to toxicity</p>	Hyperkalemia	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p><u>Adverse Event</u> Hypermagnesemia</p> <p><u>Grade 1</u>: >ULN - 3.0 mg/dL; >ULN - 1.23 mmol/L <u>N/A Grade 2</u>: <u>Grade 3</u>: >3.0 - 8.0 mg/dL; >1.23 - 3.30 mmol/L <u>Grade 4</u>: >8.0 mg/dL; >3.30 mmol/L <u>Grade 5</u>: death related to toxicity</p>	Hypermagnesemia	1, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			

Category: METABOLIC/LABORAT

<p>Adverse Event Hypernatremia</p> <p>Grade 1: >UNL - 150 mmol/L Grade 2: >150 - 155 mmol/L Grade 3: >155 - 160 mmol/L Grade 4: >160 mmol/L Grade 5: death related to toxicity</p>	Hypernatremia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hypertriglyceridemia</p> <p>Grade 1: >UNL - 2.5 x ULN Grade 2: >2.5 - 5.0 x ULN Grade 3: >5.0 - 10 x ULN Grade 4: >10 x ULN Grade 5: death related to toxicity</p>	Hypertriglyceridemia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hyperuricemia</p> <p>Grade 1: >ULN - <= 10 mg/dL; <= 0.59 mmol/L without physiologic consequences N/A Grade 2: Grade 3: >ULN - 10 mg/dL; <= 0.59 mmol/L with physiologic consequences Grade 4: >10 mg/dL; >0.59 mmol/L Grade 5: death related to toxicity</p> <p>Note: Also consider Tumor lysis syndrome, Renal failure, Creatinine, Hyperkalemia.</p>	Hyperuricemia	1, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hypocalcemia</p> <p>Grade 1: < LLN - 8.0 mg/dL; < LLN - 2.0 mmol/L Grade 2: 7.0 - <8.0 mg/dL; 1.75 - <2.0 mmol/L Grade 3: 6.0 - <7.0 mg/dL; 1.5 - <1.75 mmol/L Grade 4: <6.0 mg/dL; <1.5 mmol/L Grade 5: death related to toxicity</p>	Hypocalcemia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			

<p>Adverse Event Hypoglycemia</p> <p>Grade 1: < LLN - 55 mg/dL; < LLN - 3.0 mmol/L Grade 2: 40 - <55 mg/dL; 2.2 - <3.0 mmol/L Grade 3: 30 - <40 mg/dL; 1.7 - 2.2 mmol/L Grade 4: <30 mg/dL; <1.7 mmol/L Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hypokalemia</p> <p>Grade 1: < LLN - 3.0 mmol/L N/A Grade 2: Grade 3: 2.5 - <3.0 mmol/L Grade 4: <2.5 mmol/L Grade 5: death related to toxicity</p>	Hypokalemia	1, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hypomagnesemia</p> <p>Grade 1: < LLN - 1.2 mg/dL; < LLN - 0.5 mmol/L Grade 2: 0.9 - <1.2 mg/dL; 0.4 - <0.5 mmol/L Grade 3: 0.7 - <0.9 mg/dL; 0.3 - <0.4 mmol/L Grade 4: <0.7 mg/dL; <0.3 mmol/L Grade 5: death related to toxicity</p>	Hypomagnesemia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hyponatremia</p> <p>Grade 1: < LLN - 130 mmol/L N/A Grade 2: Grade 3: 120 - <130 mmol/L Grade 4: <120 mmol/L Grade 5: death related to toxicity</p>	Hyponatremia	1, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hypophosphatemia</p> <p>Grade 1: < LLN - 2.5 mg/dL; < LLN - 0.8 mmol/L Grade 2: >=2.0 - <2.5 mg/dL; >=0.6 - <0.8 mmol/L Grade 3: >=1.0 - <2.0 mg/dL; >=0.3 - <0.6 mmol/L Grade 4: <1.0 mg/dL; <0.3 mmol/L Grade 5: death related to toxicity</p>	Hypophosphatemia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			

Hypothyroidism is graded in the ENDOCRINE category										
<p>Adverse Event Lipase</p> <p>Grade 1: >ULN - 1.5 x ULN Grade 2: >1.5 - 2.0 x ULN Grade 3: >2.0 - 5.0 x ULN Grade 4: >5.0 x ULN Grade 5: death related to toxicity</p>										
<p>Adverse Event Metabolic/Laboratory-Other (Specify,_____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening of disabling Grade 5: death related to toxicity</p>										
MUSCULOSKELETAL										
Toxicity category and code name										
Description (reported term)										
Grade										
LLT MedDRA										
Preferred term MedDRA										
HLT										
HLGT										
Pr = Primary SOC Sec_n = Secondary SOC										
SSC1										
SSC2										
SSC3										
Arthralgia is graded in the PAIN category										
<p>Adverse Event Arthritis</p> <p>Grade 1: mild pain with inflammation, erythema or joint swelling but not interfering with function Grade 2: moderate pain with inflammation, erythema, or joint swelling interfering with function, but not interfering with activities of daily living Grade 3: severe pain with inflammation, erythema, or joint swelling and interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>										
<p>Adverse Event Muscle weakness (not due to neuropathy)</p> <p>Grade 1: asymptomatic with weakness on physical exam Grade 2: symptomatic and interfering with function, but not interfering with activities of daily living</p>										

Category: MUSCULOSKELETAL	<p>activities of daily living</p> <p>Grade 3: symptomatic and interfering with activities of daily living</p> <p>Grade 4: bedridden or disabling</p> <p>Grade 5: death related to toxicity</p>	MUSCLE WEAKNESS	4	LLT :	FL :		HLT : Neuromuscular disorders NEC	HLGT : Neuromuscular disorders	Sec ₁ : Nervous system disorders			
	<p>Adverse Event Myositis (inflammation/damage of muscle)</p> <p>Grade 1: mild pain, not interfering with function</p> <p>Grade 2: pain interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: pain interfering with function and interfering with activities of daily living</p> <p>Grade 4: bedridden or disabling</p> <p>Grade 5: death related to toxicity</p> <p>Note: Myositis implies muscle damage (i.e., elevated CPK). Also consider CPK.</p>	Myositis	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
				LLT :								
	<p>Note: Myositis implies muscle damage (i.e., elevated CPK). Also consider CPK.</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
				LLT :		HLT :	HLGT :	Sec ₁ :				
	Myalgia is graded in the PAIN category											
	<p>Adverse Event Osteonecrosis (avascular necrosis)</p> <p>Grade 1: asymptomatic and detected by imaging only</p> <p>Grade 2: symptomatic and interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: symptomatic and interfering with activities of daily living</p> <p>Grade 4: symptomatic; or disabling</p> <p>Grade 5: death related to toxicity</p>	Osteonecrosis	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
				LLT :	PT :	HLT :	HLGT :	Pr :				
	<p>Adverse Event Musculoskeletal-Other (Specify, _____)</p> <p>Grade 1: mild</p> <p>Grade 2: moderate</p> <p>Grade 3: severe</p> <p>Grade 4: life-threatening or disabling</p> <p>Grade 5: death related to toxicity</p>											

NEUROLOGY

Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
Aphasia, receptive and/or expressive, is graded under Speech impairment in the NEUROLOGY category										
<p>Adverse Event Arachnoiditis/meningismus/radiculitis</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain interfering with function, but not interfering with activities of daily living Grade 3: severe pain interfering with activities of daily living Grade 4: unable to function or perform activities of daily living; bedridden; paraplegia Grade 5: death related to toxicity</p> <p>Note: Also consider Headache, Vomiting, Fever.</p>	Arachnoiditis/meningismus/radiculitis		LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Radiculitis		LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Ataxia (incoordination)</p> <p>Grade 1: asymptomatic but abnormal on physical exam, and not interfering with function Grade 2: mild symptoms interfering with function, but not interfering with activities of daily living Grade 3: moderate symptoms interfering with activities of daily living Grade 4: bedridden or disabling Grade 5: death related to toxicity</p>	Ataxia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event CNS cerebrovascular ischemia</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: transient ischemic event or attack (TIA) Grade 4: permanent event (e.g., cerebral vascular accident) Grade 5: death related to toxicity</p>	CNS cerebrovascular ischemia		LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT :	Sec₁ :			
		3	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT :	Sec₁ :			
CNS hemorrhage/bleeding is graded in the HEMORRHAGE category										
<p>Adverse Event Cognitive disturbance/learning problems (for pediatrics)</p> <p>Grade 1: cognitive disability; not interfering with work/school performance; preservation of intelligence Grade 2: cognitive disability; interfering with work/school performance; decline of 1 SD (Standard Deviation) or loss of developmental milestones Grade 3: cognitive disability; resulting in significant impairment of work/school performance; cognitive decline > 2 SD</p>	Cognitive disturbance/learning problems	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT :	Sec₁ :			

<p>Grade 4: inability to work/frank mental retardation Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Confusion</p> <p>Grade 1: confusion or disorientation or attention deficit of brief duration; resolves spontaneously with no sequelae Grade 2: confusion or disorientation or attention deficit interfering with function, but not interfering with activities of daily living Grade 3: confusion or delirium interfering with activities of daily living Grade 4: harmful to others or self; requiring hospitalization Grade 5: death related to toxicity</p>	Confusion	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Disorientation		LLT :	PT :	HLT :	HLGT :	Pr :			
		1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :	Sec1 :		
	Delirium	3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Cranial neuropathy is graded in the NEUROLOGY category as Neuropathy-cranial</p>										
<p>Adverse Event Delusions</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present Grade 4: toxic psychosis Grade 5: death related to toxicity</p>	Delusions	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Depressed level of consciousness</p> <p>Grade 1: somnolence or sedation not interfering with function Grade 2: somnolence or sedation interfering with function, but not interfering with activities of daily living Grade 3: obtundation or stupor; difficult to arouse; interfering with activities of daily living Grade 4: coma Grade 5: death related to toxicity</p> <p>Note: Syncope (fainting) is graded in the NEUROLOGY category.</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Somnolence	1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :	Sec1 :		
		1 or 2	LLT :	PT :	HLT :	HLGT :	Pr :			
	Obtundation	3	LLT :	PT :	HLT :	HLGT :	Pr :			

	Stupor		LLT :	PT :	HLT :	HLGT :	Pr :			
		3	LLT :	PT :	HLT :	HLGT :	Pr :			
	Coma		LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Dizziness/lightheadedness</p> <p>Grade 1 : not interfering with function</p> <p>Grade 2 : interfering with function, but not interfering with activities of daily living</p> <p>Grade 3 : interfering with activities of daily living</p> <p>Grade 4 : bedridden or disabling</p> <p>Grade 5 : death related to toxicity</p>	Dizziness/lightheadedness	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT : Neurological disorders NEC	Pr : Nervous system disorders			
			1, 2, 3 or 4					LLT :		
Dysphasia, receptive and/or expressive, is graded under Speech impairment in the NEUROLOGY category										
<p>Adverse Event Extrapryramidal/involuntary movement/restlessness</p> <p>Grade 1 : mild involuntary movements not interfering with function</p> <p>Grade 2 : moderate involuntary movements interfering with function, but not interfering with activities of daily living</p> <p>Grade 3 : severe involuntary movements or torticollis interfering with activities of daily living</p> <p>Grade 4 : bedridden or disabling</p> <p>Grade 5 : death related to toxicity</p>	Extrapryramidal/involuntary movement/restlessness	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :		
						HLT :	HLGT :	Sec1 :		
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Hallucinations</p> <p>N/A Grade 1 :</p> <p>N/A Grade 2 :</p> <p>Grade 3 : present</p> <p>Grade 4 : toxic psychosis</p> <p>Grade 5 : death related to toxicity</p>	Hallucinations	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Headache is graded in the PAIN category									

Category: NEUROLOGY

<p>Adverse Event Insomnia</p> <p>Grade 1: occasional difficulty sleeping not interfering with function Grade 2: difficulty sleeping interfering with function, but not interfering with activities of daily living Grade 3: frequent difficulty sleeping, interfering with activities of daily living N/A Grade 4: Grade 5: death related to toxicity</p> <p>Note: This adverse event is graded when insomnia is related to treatment. If pain or other symptoms interfere with sleep do NOT grade as insomnia.</p>	Insomnia	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
	Difficulty sleeping		LLT :		HLT :	HLGT :	Sec :			
<p>Adverse Event Irritability (children <3 years of age)</p> <p>Grade 1: mild; easily consolable Grade 2: moderate; requiring increased attention Grade 3: severe; inconsolable N/A Grade 4: Grade 5: death related to toxicity</p>	Irritability	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Leukoencephalopathy associated with radiological findings</p> <p>Grade 1: mild increase in SAS (subarachnoid space) and/or mild ventriculomegaly; and/or small (+/- multiple) focal T2 hyperintensities, involving periventricular white matter or <1/3 of susceptible areas of cerebrum Grade 2: moderate increase in SAS; and/or moderate ventriculomegaly; and/or focal T2 hyperintensities extending into centrum ovale; or involving 1/3 to 2/3 of susceptible areas of cerebrum Grade 3: severe increase in SAS; severe ventriculomegaly; near total white matter T2 hyperintensities or diffuse low attenuation (CT); focal white matter necrosis (cystic) Grade 4: severe increase in SAS; severe ventriculomegaly; diffuse low attenuation with calcification (CT); diffuse white matter necrosis (MRI) Grade 5: death related to toxicity</p>	Leukoencephalopathy	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Memory loss</p> <p>Grade 1: memory loss not interfering with function Grade 2: memory loss interfering with function, but not interfering with activities of daily living Grade 3: memory loss interfering with activities of daily living Grade 4: amnesia Grade 5: death related to toxicity</p>	Memory loss	1, 2, 3 or 4	LLT : Memory loss	PT : Memory impairment	HLT :	HLGT :	Pr :			
					HLT :	HLGT :	Sec :			
		4	LLT :	PT :	HLT :	HLGT :	Pr :	Sec :		
<p>Adverse Event Mood alteration-anxiety, agitation</p>	Mood alteration-anxiety, agitation	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			

<p>Grade 1: mild mood alteration not interfering with function</p> <p>Grade 2: moderate mood alteration interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: severe mood alteration interfering with activities of daily living</p> <p>Grade 4: suicidal ideation or danger to self</p> <p>Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Mood alteration-depression</p> <p>Grade 1: mild mood alteration not interfering with function</p> <p>Grade 2: moderate mood alteration interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: severe mood alteration interfering with activities of daily living</p> <p>Grade 4: suicidal ideation or danger to self</p> <p>Grade 5: death related to toxicity</p>	Mood alteration-depression	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Mood alteration-euphoria</p> <p>Grade 1: mild mood alteration not interfering with function</p> <p>Grade 2: moderate mood alteration interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: severe mood alteration interfering with activities of daily living</p> <p>Grade 4: danger to self</p> <p>Grade 5: death related to toxicity</p>	Mood alteration-euphoria	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Neuropathic pain is graded in the PAIN category.</p>										
<p>Adverse Event Neuropathy - cranial</p> <p>N/A Grade 1:</p> <p>Grade 2: present, not interfering with activities of daily living</p> <p>Grade 3: present, interfering with activities of daily living</p> <p>Grade 4: life-threatening, disabling</p> <p>Grade 5: death related to toxicity</p> <p>Attribution Code Unrelated Unlikely Possible Probable Definite</p>	Neuropathy - cranial	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Neuropathy - motor</p> <p>Grade 1: subjective weakness but no objective findings</p> <p>Grade 2: mild objective weakness interfering with function, but not interfering with activities of daily living</p>	Neuropathy - motor	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			

<p>interfering with activities of daily living Grade 3: objective weakness interfering with activities of daily living Grade 4: paralysis Grade 5: death related to toxicity</p>		4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Neuropathy-sensory Grade 1: loss of deep tendon reflexes or paresthesia (including tingling) but not interfering with function Grade 2: objective sensory loss or paresthesia (including tingling), interfering with function, but not interfering with activities of daily living Grade 3: sensory loss or paresthesia interfering with activities of daily living Grade 4: permanent sensory loss that interferes with function Grade 5: death related to toxicity</p>	Neuropathy-sensory	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Loss of deep tendon reflexes	1	LLT :	PT :	HLT :	HLGT :	Pr :			
	Paresthesia		LLT :	PT :	HLT :	HLGT :	Pr :			
		1 or 2	LLT :							
<p>Adverse Event Nystagmus Grade 1: present N/A Grade 2: N/A Grade 3: N/A Grade 4: N/A Grade 5: Note: Also consider Vision-double vision.</p>	Nystagmus	1	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Personality/behavioral Grade 1: change, but not disruptive to patient or family Grade 2: disruptive to patient or family Grade 3: disruptive to patient and family; requiring mental health intervention Grade 4: harmful to others or self; requiring hospitalization Grade 5: death related to toxicity</p>	Personality/behavioral	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Pyramidal tract dysfunction (e.g., increased tone, hyperreflexia, positive Babinski, decreased fine motor coordination) Grade 1: asymptomatic with abnormality on physical examination Grade 2: symptomatic or interfering with function but not interfering with activities of daily living Grade 3: interfering with activities of daily living Grade 4: bedridden or disabling; paralysis Grade 5: death related to toxicity</p>	Pyramidal tract dysfunction	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT :	Pr :			

		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
<p>Adverse Event Seizure(s)</p> <p>N/A Grade 1: Grade 2: seizure(s) self-limited and consciousness is preserved Grade 3: seizure(s) in which consciousness is altered Grade 4: seizures of any type which are prolonged, repetitive, or difficult to control (e.g., status epilepticus, intractable epilepsy) Grade 5: death related to toxicity</p>	Seizure	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
		4	LLT :	PT :	HLT :	HLGT :	Pr :				
		4	LLT :	PT :	HLT :	HLGT :	Pr :				
<p>Adverse Event Speech impairment (e.g., dysphasia or aphasia)</p> <p>N/A Grade 1: Grade 2: awareness of receptive or expressive dysphasia, not impairing ability to communicate Grade 3: receptive or expressive dysphasia, impairing ability to communicate Grade 4: inability to communicate Grade 5: death related to toxicity</p>	Speech impairment	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
		2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
		2, 3 or 4	LLT :		HLT :	HLGT :	Pr :				
				LLT :		HLT :	HLGT :	Sec₁ :			
		Aphasia	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			2, 3 or 4	LLT :		HLT :	HLGT :	Pr :			
		2, 3 or 4	LLT :	HLT :		HLGT :	Sec₁ :				
<p>Adverse Event Syncope (fainting)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present N/A Grade 4: Grade 5: death related to toxicity</p> <p>Note: Also consider CARDIOVASCULAR (ARRHYTHMIA), Vasovagal episode, CNS cerebrovascular ischemia.</p>	Syncope	3	LLT :	PT :	HLT :	HLGT : Neurological disorders NEC	Pr : Nervous system disorders	SSC₁ : ARREST (CARDIAC)			
			3						LLT :		

<p>Adverse Event Tremor</p> <p>Grade 1: mild and brief or intermittent but not interfering with function Grade 2: moderate tremor interfering with function, but not interfering with activities of daily living Grade 3: severe tremor interfering with activities of daily living N/A Grade 4: Grade 5: death related to toxicity</p>	Tremor	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Vertigo</p> <p>Grade 1: not interfering with function Grade 2: interfering with function, but not interfering with activities of daily living Grade 3: interfering with activities of daily living Grade 4: bedridden or disabling Grade 5: death related to toxicity</p>	Vertigo	1, 2, 3 or 4	LLT : Vertigo	PT : Vertigo	HLT : Inner ear signs and symptoms	HLGT : Inner ear and VIIIth cranial nerve disorders	Pr : Ear and labyrinth disorders			
<p>Adverse Event Neurology-Other (Specify, _____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>					HLT : Vertigos NEC	HLGT : Neurological disorders NEC	Sec₁ : Nervous system disorders			
OCULAR/VISUAL										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Cataract</p> <p>Grade 1: asymptomatic Grade 2: symptomatic, partial visual loss Grade 3: symptomatic, visual loss requiring treatment or interfering with function N/A Grade 4: Grade 5: death related to toxicity</p>	Cataract	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			

<p><u>Adverse Event</u> Conjunctivitis</p> <p><u>Grade 1</u>: abnormal ophthalmologic changes, but asymptomatic or symptomatic without visual impairment (i.e., pain and irritation)</p> <p><u>Grade 2</u>: symptomatic and interfering with function, but not interfering with activities of daily living</p> <p><u>Grade 3</u>: symptomatic and interfering with activities of daily living</p> <p><u>N/A Grade 4</u>:</p> <p><u>Grade 5</u>: death related to toxicity</p>		1, 2 or 3	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :				
<p><u>Adverse Event</u> Dry eye</p> <p><u>Grade 1</u>: mild, not requiring treatment</p> <p><u>Grade 2</u>: moderate or requiring artificial tears</p> <p><u>N/A Grade 3</u>:</p> <p><u>N/A Grade 4</u>:</p> <p><u>N/A Grade 5</u>:</p>	Dry eye	1 or 2	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :				
<p><u>Adverse Event</u> Glaucoma</p> <p><u>Grade 1</u>: increase in intraocular pressure but no visual loss</p> <p><u>Grade 2</u>: increase in intraocular pressure with retinal changes</p> <p><u>Grade 3</u>: visual impairment</p> <p><u>Grade 4</u>: unilateral or bilateral loss of vision (blindness)</p> <p><u>Grade 5</u>: death related to toxicity</p>	Glaucoma	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :				
		1 or 2	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :				
		3	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :				
		4	<u>LLT</u> :		<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₁</u> :				
	Bilateral loss of vision		<u>LLT</u> :		<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
		4	<u>LLT</u> :		<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Sec₁</u> :			
	Keratitis	1, 2, 3 or 4	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :				

Category: OCULAR/VISUAL

<p>Adverse Event Keratitis (corneal inflammation/corneal ulceration)</p> <p>Grade 1: abnormal ophthalmologic changes but asymptomatic or symptomatic without visual impairment (i.e., pain and irritation)</p> <p>Grade 2: symptomatic and interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: symptomatic and interfering with activities of daily living</p> <p>Grade 4: unilateral or bilateral loss of vision (blindness)</p> <p>Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PL :	HLT :	HLGT :	Pr :			
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
		4	LLT :		HLT :	HLGT :	Pr : Eye disorders			
		4	LLT :	PT :						
	Blindness	4	LLT :		HLT : Neurologic visual problems NEC	HLGT : Neurological disorders of the eye	Sec : Nervous system disorders			
<p>Adverse Event Tearing (watery eyes)</p> <p>Grade 1: mild: not interfering with function</p> <p>Grade 2: moderate: interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: interfering with activities of daily living</p> <p>N/A Grade 4:</p> <p>Grade 5: death related to toxicity</p>		1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2 or 3	LLT :							
<p>Adverse Event Vision-blurred vision</p> <p>N/A Grade 1:</p> <p>Grade 2: symptomatic and interfering with function, but not interfering with activities of daily living</p> <p>Grade 3: symptomatic and interfering with activities of daily living</p> <p>N/A Grade 4:</p> <p>Grade 5: death related to toxicity</p>	Blurred vision	2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
					HLT :	HLGT :	Sec :			
<p>Adverse Event Vision-double vision (diplopia)</p> <p>N/A Grade 1:</p> <p>Grade 2: symptomatic and interfering with function, but not interfering with</p>	Double vision		LLT :	PT :	HLT :	HLGT :	Pr :			

<p>activities of daily living <u>Grade 3</u>: symptomatic and interfering with activities of daily living <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>		2 or 3	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Sec</u> ₁ :			
<p><u>Adverse Event</u> Vision-flashing lights/floaters <u>Grade 1</u>: mild, not interfering with function <u>Grade 2</u>: symptomatic and interfering with function, but not interfering with activities of daily living <u>Grade 3</u>: symptomatic and interfering with activities of daily living <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>	Flashing lights		<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p><u>Adverse Event</u> Vision-night blindness (nyctalopia) <u>Grade 1</u>: abnormal electro-retinography but asymptomatic <u>Grade 2</u>: symptomatic and interfering with function, but not interfering with activities of daily living <u>Grade 3</u>: symptomatic and interfering with activities of daily living <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>	Nyctalopia	1, 2 or 3	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p><u>Adverse Event</u> Vision-photophobia <u>N/A Grade 1</u>: <u>Grade 2</u>: symptomatic and interfering with function, but not interfering with activities of daily living <u>Grade 3</u>: symptomatic and interfering with activities of daily living <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>			<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
<p><u>Adverse Event</u> Ocular/Visual-Other (Specify, ___) <u>Grade 1</u>: mild <u>Grade 2</u>: moderate <u>Grade 3</u>: severe <u>Grade 4</u>: unilateral or bilateral loss of vision (blindness) <u>Grade 5</u>: death related to toxicity</p>										
PAIN										

Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Abdominal pain or cramping</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Abdominal pain	1, 2, 3 or 4	LLT :							
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
<p>Adverse Event Arthralgia (joint pain)</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Arthralgia	1, 2, 3 or 4	LLT :							
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Arthritis (joint pain with clinical signs of inflammation) is graded in the MUSCULOSKELETAL category</p>										
<p>Adverse Event Bone pain</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Bone pain	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC : PAIN		
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :	SSC : PAIN	
<p>Adverse Event Chest pain (non-cardiac and non-pleuritic)</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living</p>		1, 2, 3 or 4	LLT :							
		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :	SSC : PAIN	

<p>Grade 4: disabling Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :							
<p>Adverse Event Dysmenorrhea</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Dysmenorrhea	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Dyspareunia</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain interfering with sexual activity Grade 3: severe pain preventing sexual activity N/A Grade 4: Grade 5: death related to toxicity</p>	Dyspareunia	1, 2 or 3	LLT : Dyspareunia	PT : Dyspareunia NOS	HLT :	HLGT :	Pr :			
<p>Dysuria is graded in the RENAL/GENITOURINARY category</p>										
<p>Adverse Event Earache (otalgia)</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Earache	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
<p>Adverse Event Headache</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Headache	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC : PAIN		

IN

Category: PA	<p>Adverse Event Hepatic pain</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Hepatic pain	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : PAIN			
	<p>Adverse Event Myalgia (muscle pain)</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Myalgia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :				
	<p>Adverse Event Neuropathic pain (e.g., jaw pain, neurologic pain, phantom limb pain, post-infectious neuralgia, or painful neuropathies)</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Neuropathic pain	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : PAIN			
					LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : PAIN		
				1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : PAIN		
		Neuralgia	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : PAIN			
	<p>Adverse Event Pain due to radiation</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>		1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ : PAIN			
						HLT :	HLGT :	Sec₁ :				

<p>Adverse Event Pelvic pain</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Pelvic pain	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
<p>Adverse Event Pleuritic pain</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Pleuritic pain		LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
<p>Adverse Event Rectal or perirectal pain (proctalgia)</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Rectal pain	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC :		
		1, 2, 3 or 4	LLT :							
		1, 2, 3 or 4	LLT :							
<p>Adverse Event Tumor pain (onset or exacerbation of tumor pain due to treatment)</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain: pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain: pain or analgesics severely interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p>	Tumor pain	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC : PAIN		
<p>Tumor flair is graded in the SYNDROME category</p>										

<p>Adverse Event Pain-Other (Specify,____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: disabling Grade 5: death related to toxicity</p>										
PULMONARY										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Adult respiratory distress syndrome (ARDS)</p> <p>N/A Grade 1: N/A Grade 2: N/A Grade 3: Grade 4: present Grade 5: death related to toxicity</p>	Adult respiratory distress syndrome	4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Apnea</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present Grade 4: requiring intubation Grade 5: death related to toxicity</p>	Apnea	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Carbon monoxide diffusion capacity (DL(co))</p> <p>Grade 1: >= 75 - <90% of pretreatment or normal value Grade 2: >= 50 - <75% of pretreatment or normal value Grade 3: >= 25 - <50% of pretreatment or normal value Grade 4: <25% of pretreatment or normal value Grade 5: death related to toxicity</p>	Carbon monoxide diffusion capacity (DL(co))	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Cough</p> <p>Grade 1: mild, relieved by non-prescription medication Grade 2: requiring narcotic antitussive Grade 3: severe cough or coughing spasms, poorly controlled or unresponsive to treatment N/A Grade 4: Grade 5: death related to toxicity</p>	Cough	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Dyspnea (shortness of breath)</p>	Dyspnea	2, 3 or 4	LLT :	PT :	HLT :	HLGT : Respiratory disorders NEC	Pr : Respiratory, thoracic and mediastinal disorders	SSC1 : ANAPHYL AXI	SSC2 : CARDIAC	

			2, 3 or 4	LLT :	PL :	HLT :	HLGT : Cardiac disorder signs and symptoms	Sec₁ : Cardiac disorders	ANAPHYLACTOIDS	CARDIAC ISCHAEMIA	
	N/A Grade 1 : Grade 2 : dyspnea on exertion Grade 3 : dyspnea at normal level of activity Grade 4 : dyspnea at rest or requiring ventilator support Grade 5 : death related to toxicity			LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :		
				LLT :		HLT :	HLGT :	Sec₁ :			
	Adverse Event FEV (1) Grade 1 : >= 75 - <90% of pretreatment or normal value Grade 2 : >= 50 - <75% of pretreatment or normal value Grade 3 : >= 25 - <50% of pretreatment or normal value Grade 4 : <25% of pretreatment or normal value Grade 5 : death related to toxicity	FEV (1)	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			1, 2 or 3	LLT :		HLT :	HLGT :	Pr : Gastrointestinal disorders			
	Adverse Event Hiccoughs (hiccups, singultus) Grade 1 : mild, not requiring treatment Grade 2 : moderate, requiring treatment Grade 3 : severe, prolonged and refractory to treatment N/A Grade 4 : Grade 5 : death related to toxicity		1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Sec₁ :			
			1, 2 or 3	LLT :		HLT :	HLGT :	Sec₁ :			
	Adverse Event Hypoxia N/A Grade 1 : Grade 2 : decreased O2 saturation with exercise Grade 3 : decreased O2 saturation at rest, requiring supplemental oxygen Grade 4 : decreased O2 saturation, requiring pressure support (CPAP) or assisted ventilation Grade 5 : death related to toxicity	Hypoxia	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Adverse Event Pleural effusion (non-malignant) Grade 1 : asymptomatic and not requiring treatment Grade 2 : symptomatic, requiring diuretics Grade 3 : symptomatic, requiring O2 or therapeutic thoracentesis Grade 4 : life-threatening (e.g., requiring intubation) Grade 5 : death related to toxicity	Pleural effusion (non-malignant)	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Pleuritic pain is graded in the PAIN category										

<p>Adverse Event Pneumonitis/pulmonary infiltrates</p> <p>Grade 1: radiographic changes but asymptomatic or symptoms not requiring steroids</p> <p>Grade 2: radiographic changes and requiring steroids or diuretics</p> <p>Grade 3: radiographic changes and requiring oxygen</p> <p>Grade 4: radiographic changes and requiring assisted ventilation</p> <p>Grade 5: death related to toxicity</p>	<p>Pneumonitis/pulmonary infiltrates</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
		<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Adverse Event Pneumothorax</p> <p>Grade 1: no intervention required</p> <p>Grade 2: chest tube required</p> <p>Grade 3: sclerosis or surgery required</p> <p>Grade 4: life-threatening</p> <p>Grade 5: death related to toxicity</p>	<p>Pneumothorax</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Pulmonary embolism is graded as Thrombosis/embolism in the CARDIOVASCULAR (GENERAL) category</p>										
<p>Adverse Event Pulmonary fibrosis</p> <p>Grade 1: radiographic changes, but asymptomatic or symptoms not requiring steroids</p> <p>Grade 2: requiring steroids or diuretics</p> <p>Grade 3: requiring oxygen</p> <p>Grade 4: requiring assisted ventilation</p> <p>Grade 5: death related to toxicity</p> <p>Note: Radiation-related pulmonary fibrosis is graded in the RTOG/EORTC Late Radiation Morbidity Scoring Scheme-Lung (see Appendix IV).</p>	<p>Pulmonary fibrosis</p>	<p>1, 2, 3 or 4</p>	<p>LLT : Pulmonary fibrosis</p>	<p>PT : Pulmonary fibrosis</p>	<p>HLT : Parenchymal lung disorders NEC</p>	<p>HLGT : Lower respiratory tract disorders (excl obstruction and infection)</p>	<p>Pr : Respiratory, thoracic and mediastinal disorders</p>			
<p>Adverse Event Voice changes/stridor/larynx (e.g., hoarseness, loss of voice, laryngitis)</p> <p>Grade 1: mild or intermittent hoarseness</p> <p>Grade 2: persistent hoarseness, but able to vocalize; may have mild to moderate edema</p> <p>Grade 3: whispered speech, not able to vocalize; may have marked edema</p> <p>Grade 4: marked dyspnea/stridor requiring tracheostomy or intubation</p> <p>Grade 5: death related to toxicity</p>	<p>Voice changes/stridor/larynx (e.g., hoarseness, loss of voice, laryngitis)</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Grade 2: persistent hoarseness, but able to vocalize; may have mild to moderate edema</p>		<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>	<p>SSC: ANAPHYLAXIS</p>		
<p>Grade 3: whispered speech, not able to vocalize; may have marked edema</p> <p>Grade 4: marked dyspnea/stridor requiring tracheostomy or intubation</p> <p>Grade 5: death related to toxicity</p>		<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			
<p>Note: Cough from radiation is graded as cough in the PULMONARY category. Radiation-related hemoptysis from larynx/pharynx is graded as Grade 4 Mucositis due to radiation in the GASTROINTESTINAL category. Radiation-related hemoptysis from the thoracic cavity is graded as Grade 4 Hemoptysis in the HEMORRHAGE category.</p>	<p>Loss of voice</p>	<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>	<p>Sec:</p>		
		<p>1, 2, 3 or 4</p>	<p>LLT :</p>	<p>PT :</p>	<p>HLT :</p>	<p>HLGT :</p>	<p>Pr :</p>			

<p>Adverse Event Pulmonary-Other (Specify,____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
RENAL/GENITOURINARY										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
<p>Adverse Event Bladder spasms</p> <p>Grade 1: mild symptoms, not requiring intervention Grade 2: symptoms requiring antispasmodic Grade 3: severe symptoms requiring narcotic N/A Grade 4: Grade 5: death related to toxicity</p>	Bladder spasms	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Creatinine</p> <p>Grade 1: > ULN - 1.5 x ULN Grade 2: > 1.5 - 3.0 x ULN Grade 3: > 3.0 - 6.0 x ULN Grade 4: > 6.0 x ULN Grade 5: death related to toxicity</p> <p>Note: Adjust to age-appropriate levels for pediatric patients.</p>	Creatinine	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Dysuria (painful urination)</p> <p>Grade 1: mild symptoms requiring no intervention Grade 2: symptoms relieved with therapy Grade 3: symptoms not relieved despite therapy N/A Grade 4: Grade 5: death related to toxicity</p>		1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Fistula or GU fistula (e.g., vaginal, vesicovaginal)</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: requiring intervention Grade 4: requiring surgery Grade 5: death related to toxicity</p>	Fistula or GU fistula (e.g., vaginal, vesicovaginal)	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			Sec₁ :

ity: RENAL/GENITOURINARY

<p>Adverse Event Hemoglobinuria</p> <p>Grade 1: present N/A Grade 2: N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Hemoglobinuria	1	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Hematuria (in the absence of vaginal bleeding) is graded in the HEMORRHAGE category</p>										
<p>Adverse Event Incontinence</p> <p>Grade 1: with coughing, sneezing, etc. Grade 2: spontaneous, some control Grade 3: no control (in the absence of fistula) N/A Grade 4: Grade 5: death related to toxicity</p>	Incontinence	1, 2 or 3	LLT : Incontinence	PT : Incontinence NOS	HLT :	HLGT :	Pr :			
					HLT :	HLGT :	Sec₁ :			
	HLT : Autonomic nervous system disorders	HLGT : Neuromuscular disorders	Sec₂ : Nervous system disorders							
	LLT :	PT :	HLT :	HLGT :	Pr :	Sec₁ :				
<p>Adverse Event Operative injury to bladder and/or ureter</p> <p>N/A Grade 1: Grade 2: injury of bladder with primary repair Grade 3: sepsis, fistula, or obstruction requiring secondary surgery; loss of one kidney; injury requiring anastomosis or re-implantation Grade 4: septic obstruction of both kidneys or vesicovaginal fistula requiring diversion Grade 5: death related to toxicity</p>	Operative injury to bladder and/or ureter	2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Proteinuria</p> <p>Grade 1: 1+ or 0.15 - 1.0 g/24 hour Grade 2: 2+ to 3+ or 1.0 - 3.5 g/24 hour Grade 3: 4+ or > 3.5 g/24 hour Grade 4: nephrotic syndrome Grade 5: death related to toxicity</p> <p>Note: If there is an inconsistency between absolute value and dip stick reading, use the absolute value for grading.</p>	Proteinuria	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :		
<p>Adverse Event Renal failure</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: requiring dialysis, but reversible Grade 4: requiring dialysis and irreversible Grade 5: death related to toxicity</p>	Renal failure	3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			SSC₁ : SECONDARY IMMUNOCOM PROMISED STATE

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<p>Adverse Event Ureteral obstruction</p> <p>Grade 1: unilateral, not requiring surgery N/A Grade 2: Grade 3: bilateral, not requiring surgery Grade 4: stent, nephrostomy tube, or surgery Grade 5: death related to toxicity</p>	Ureteral obstruction		LLT :	PT :	HLT :	HLGT :	Pr :			
		4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Ureteral stent	4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Urinary electrolyte wasting (e.g., Fanconi's syndrome, renal tubular acidosis)</p> <p>Grade 1: asymptomatic, not requiring treatment Grade 2: mild, reversible and manageable with oral replacement Grade 3: reversible but requiring IV replacement Grade 4: irreversible, requiring continued replacement Grade 5: death related to toxicity</p> <p>Note: Also consider Acidosis, Bicarbonate, Hypocalcemia, Hypophosphatemia.</p>	Urinary electrolyte wasting	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Fanconi's syndrome	1, 2, 3 or 4	LLT : Fanconi syndrome	PT : Fanconi syndrome acquired	HLT :	HLGT :	Pr :			
					HLT :	HLGT :	Sec₁ :			
				LLT :	PT :	HLT :	HLGT :	Pr :		
<p>Adverse Event Urinary frequency/urgency</p> <p>Grade 1: increase in frequency or nocturia up to 2 x normal Grade 2: increase > 2 x normal but < hourly Grade 3: hourly or more with urgency, or requiring catheter N/A Grade 4: Grade 5: death related to toxicity</p>	Urinary frequency/urgency	1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
		1, 2 or 3	LLT :							
		1, 2 or 3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Urinary retention</p> <p>Grade 1: hesitancy or dribbling, but no significant residual urine; retention occurring during the immediate postoperative period Grade 2: hesitancy requiring medication or occasional in/out catheterization (<4 x per week), or operative bladder atony requiring indwelling catheter beyond immediate postoperative period but for <6 weeks Grade 3: requiring frequent in/out catheterization (>= 4 x per week) or urological intervention (e.g., TURP, suprapubic tube, urethrotomy) Grade 4: bladder rupture Grade 5: death related to toxicity</p>	Urinary retention	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT : Injuries NEC	Pr : injury, poisoning and procedural complications			
			LLT :	PT :	HLT :	HLGT : Bladder and bladder neck disorders (excl calculi)	Sec₁ : Renal and urinary disorders			

<p>Adverse Event Urine color change (not related to other dietary or physiologic cause e.g., bilirubin, concentrated urine, hematuria)</p> <p>Grade 1: asymptomatic, change in urine color N/A Grade 2: N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>	Urine color change	1	LLT :	PT :	HLT :	HLGT :	Pr :			
Vaginal bleeding is graded in the HEMORRHAGE category										
<p>Adverse Event Vaginitis (not due to infection)</p> <p>Grade 1: mild, not requiring treatment Grade 2: moderate, relieved with treatment Grade 3: severe, not relieved with treatment, or ulceration not requiring surgery Grade 4: ulceration requiring surgery Grade 5: death related to toxicity</p>	Vaginitis	1, 2, 3 or 4	LLT : Vaginitis	PT : Vaginitis	HLT : Female reproductive tract infections	HLGT : Infections - pathogen class unspecified	Pr : Infections and infestations			
					HLT : Vaginal and vulval infections and inflammations	HLGT : Female reproductive tract infections and inflammations	Sec_n : Reproductive system and breast disorders			
<p>Adverse Event Renal/Genitourinary-Other (Specify,____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
SECONDARY MALIGNANCY										
<p>Toxicity category and code name</p>	<p>Description (reported term)</p>	<p>Grade</p>	<p>LLT MedDRA</p>	<p>Preferred term MedDRA</p>	<p>HLT</p>	<p>HLGT</p>	<p>Pr = Primary SOC Sec_n = Secondary SOC</p>	<p>SSC1</p>	<p>SSC2</p>	<p>SSC3</p>
<p>Category: SECONDARY MALIGNANCY</p> <p>Adverse Event Secondary Malignancy-Other (Specify,____) excludes metastasis from initial primary</p> <p>N/A Grade 1: N/A Grade 2: N/A Grade 3: Grade 4: present N/A Grade 5:</p>										
SEXUAL/REPRODUCTIVE FUNCTION										

	Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
	Dyspareunia is graded in the PAIN category										
	Dysmenorrhea is graded in the PAIN category										
: SEXUAL/REPRODUCTIVE FUNCTION	<p><u>Adverse Event</u> Erectile impotence</p> <p><u>Grade 1</u>: mild (erections impaired but satisfactory) <u>Grade 2</u>: moderate (erections impaired, unsatisfactory for intercourse) <u>Grade 3</u>: no erections <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>			<u>LLT</u> :	<u>PT</u> :	<p><u>HLT</u> : Erection and ejaculation conditions and disorders</p> <p><u>HLT</u> : Sexual arousal disorders</p>	<p><u>HLGT</u> : Sexual function and fertility disorders</p> <p><u>HLGT</u> : Sexual function and fertility disorders</p>	<p><u>Pr</u> : Reproductive system and breast disorders</p> <p><u>Sec₁</u> : Psychiatric disorders</p>			
	<p><u>Adverse Event</u> Female sterility</p> <p><u>N/A Grade 1</u>: <u>N/A Grade 2</u>: <u>Grade 3</u>: sterile <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>	Female sterility	3	<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			
	Feminization of male is graded in the ENDOCRINE category										
	<p><u>Adverse Event</u> Irregular menses (change from baseline)</p> <p><u>Grade 1</u>: occasionally irregular or lengthened interval, but continuing menstrual cycles <u>Grade 2</u>: very irregular, but continuing menstrual cycles <u>Grade 3</u>: persistent amenorrhea <u>N/A Grade 4</u>: <u>Grade 5</u>: death related to toxicity</p>	Irregular menses	1, 2 or 3	<u>LLT</u> : Menses irregular	<u>PT</u> : Menstruation irregular	<p><u>HLT</u> :</p> <p><u>HLT</u> :</p>	<p><u>HLGT</u> :</p> <p><u>HLGT</u> :</p>	<p><u>Pr</u> :</p> <p><u>Sec₁</u> :</p>			
				<u>LLT</u> :	<u>PT</u> :	<p><u>HLT</u> :</p> <p><u>HLT</u> :</p>	<p><u>HLGT</u> :</p> <p><u>HLGT</u> :</p>	<p><u>Pr</u> :</p> <p><u>Sec₁</u> :</p>			
	<p><u>Adverse Event</u> Libido</p> <p><u>Grade 1</u>: decrease in interest <u>Grade 2</u>: severe loss of interest <u>N/A Grade 3</u>: <u>N/A Grade 4</u>: <u>N/A Grade 5</u>:</p>		1 or 2	<u>LLT</u> :	<u>PT</u> :	<p><u>HLT</u> : Sexual desire disorders</p> <p><u>HLT</u> : Sexual function and fertility disorders NEC</p>	<p><u>HLGT</u> : Sexual dysfunctions, disturbances and gender identity disorders</p> <p><u>HLGT</u> : Sexual function and fertility disorders</p>	<p><u>Pr</u> : Psychiatric disorders</p> <p><u>Sec₁</u> : Reproductive system and breast disorders</p>			
	<p><u>Adverse Event</u> Male infertility</p>			<u>LLT</u> :	<u>PT</u> :	<u>HLT</u> :	<u>HLGT</u> :	<u>Pr</u> :			

Category:	<p>N/A Grade 1: Grade 2: oligospermia (low sperm count) Grade 3: azoospermia (no sperm) N/A Grade 4: Grade 5: death related to toxicity</p>			LLT :							
				LLT :	PT :	HLT :	HLGT :	Pr :			
	Masculinization of female is graded in the ENDOCRINE category										
	<p>Adverse Event Vaginal dryness</p> <p>Vaginal dryness 1 or 2</p> <p>Grade 1: mild Grade 2: requiring treatment and/or interfering with sexual function, dyspareunia N/A Grade 3: N/A Grade 4: N/A Grade 5:</p>			LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :	PT :	HLT :	HLGT :	Pr :				
					HLT :	HLGT :	Sec₁ :				
<p>Adverse Event Sexual/Reproductive Function-Other (Specify,____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: disabling Grade 5: death related to toxicity</p>											
SYNDROMES (not included in previous categories)											
	Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3
	Acute vascular leak syndrome is graded in the CARDIOVASCULAR (GENERAL) category	Acute vascular leak syndrome	2, 3 or 4	LLT :	PT :	HLT : HLT :	HLGT : HLGT :	Pr : Sec₁ :			
	ARDS (Adult Respiratory Distress Syndrome) is graded in the PULMONARY category		4	LLT :	PT :	HLT :	HLGT :	Pr :			
	Autoimmune reactions are graded in the ALLERGY/IMMUNOLOGY category		1, 2, 3, or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
	DIC (disseminated intravascular coagulation) is graded in the COAGULATION category	DIC	3 or 4	LLT : DIC	PT : Disseminated intravascular coagulation	HLT : HLT :	HLGT : HLGT :	Pr : Sec₁ :			

Category: SYNDROMES

Category: SYNDROMES	Fanconi's syndrome is graded as Urinary electrolyte wasting in the RENAL/GENITOURINARY category	Fanconi's syndrome	1, 2, 3 or 4	LLT : Fanconi syndrome	PT : Fanconi syndrome acquired	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			
	Renal tubular acidosis is graded as Urinary electrolyte wasting in the RENAL/GENITOURINARY category			LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			
	Stevens-Johnson syndrome (erythema multiforme) is graded in the DERMATOLOGY/SKIN category		2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			
	SIADH (syndrome of inappropriate antidiuretic hormone) is graded in the ENDOCRINE category		3	If you need to set a complete mapping version (CTC versions 2.0 and/or 3.0 - MedDRA), please contact Pharmadhoc 112, rue Olivier de Serres 75015 Paris Tel: +33 (0)8 71 76 88 17 Tel: +33 (0)6 73 51 37 96 or send an email to info@pharmadhoc.com							
	Thrombotic microangiopathy (e.g., thrombotic thrombocytopenic purpura/TTP or hemolytic uremic syndrom/HUS) is graded in the COAGULATION category			LLT :	PT :	HLT :	HLGT :	Pr :			
						HLT :	HLGT :	Sec₁ :			
			3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :	SSC₁ :		
				LLT :		HLT :	HLGT :	Sec₂ :			
			3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
			LLT :		HLT :	HLGT :	Sec₁ :				

<p>Adverse Event Tumor flare</p> <p>Grade 1: mild pain not interfering with function Grade 2: moderate pain; pain or analgesics interfering with function, but not interfering with activities of daily living Grade 3: severe pain; pain or analgesics interfering with function and interfering with activities of daily living Grade 4: disabling Grade 5: death related to toxicity</p> <p>Note: Tumor Flare is characterized by a constellation of symptoms and signs in direct relation to initiation of therapy (e.g., anti-estrogens/androgens or additional hormones). The symptoms/signs include tumor pain, inflammation of visible tumor, hypercalcemia, diffuse bone pain, and other electrolyte disturbances. Also consider Hypercalcemia.</p> <p>Attribution Code Unrelated Unlikely Possible Probable Definite</p>	Tumor flare	1, 2, 3 or 4	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Adverse Event Tumor lysis syndrome</p> <p>N/A Grade 1: N/A Grade 2: Grade 3: present N/A Grade 4: Grade 5: death related to toxicity</p> <p>Note: Also consider Hyperkalemia, Creatinine.</p>	Tumor lysis syndrome	3	LLT :	PT :	HLT :	HLGT :	Pr :			
<p>Urinary electrolyte wasting (e.g., Fanconi's syndrome, renal tubular acidosis) is graded under the RENAL/GENITOURINARY category</p>										
<p>Adverse Event Syndromes-Other (Specify,____)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening or disabling Grade 5: death related to toxicity</p>										
Appendix IV RTOG/EORTC Late Radiation Morbidity Scoring Scheme (Use for adverse events occurring greater than 90 days after radiation therapy.)										
Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec_n = Secondary SOC	SSC1	SSC2	SSC3

later than 90 days after radiation therapy.)	<p>Adverse Event Bladder- Late RT Morbidity Scoring</p> <p>Grade 1: Slight epithelial atrophy/minor telangiectasia (microscopic hematuria)</p> <p>Grade 2: Moderate frequency/generalized telangiectasia/intermittent macroscopic hematuria</p> <p>Grade 3: Severe frequency and dysuria/severe generalized telangiectasia (often with petechiae); frequent hematuria; reduction in bladder capacity (<150mL)</p> <p>Grade 4: Necrosis/contracted bladder (capacity <100 mL) severe hemorrhagic cystitis</p> <p>Grade 5: death related to adverse event</p>									
	<p>Adverse Event Bone - Late RT Morbidity Scoring</p> <p>Grade 1: Asymptomatic; no growth retardation; reduced bone density</p> <p>Grade 2: Moderate pain or tenderness; growth retardation; irregular bone sclerosis</p> <p>Grade 3: Severe pain or tenderness; complete arrest of bone growth; dense bone sclerosis</p> <p>Grade 4: Necrosis/spontaneous fracture</p> <p>Grade 5: death related to adverse event</p>									
	<p>Adverse Event Brain- Late RT Morbidity Scoring</p> <p>Grade 1: Mild headache; slight lethargy</p> <p>Grade 2: Moderate headache; great lethargy</p> <p>Grade 3: Severe headaches; severe CNS dysfunction (partial loss of power or dyskinesia)</p> <p>Grade 4: Seizures or paralysis; coma</p> <p>Grade 5: death related to adverse event</p>									
	<p>Adverse Event Esophagus- Late RT Morbidity Scoring</p> <p>Grade 1: Mild fibrosis; slight difficulty in swallowing solids; no pain on swallowing</p> <p>Grade 2: Unable to take solid food normally; swallowing semi-solid food; dilation may be indicated</p> <p>Grade 3: Severe fibrosis; able to swallow only liquids; may have pain on swallowing; dilation required</p> <p>Grade 4: Necrosis/perforation; fistula</p> <p>Grade 5: death related to adverse event</p>									

Appendix IV RTOG/EORTC Late Radiation Morbidity Scoring Scheme (Use for adverse events occurring gr

<p>Adverse Event Eye- Late RT Morbidity Scoring</p> <p>Grade 1: Asymptomatic cataract; minor corneal ulceration or keratitis Grade 2: Symptomatic cataract; moderate corneal ulceration; minor retinopathy or glaucoma Grade 3: Severe keratitis; severe retinopathy or detachment; severe glaucoma Grade 4: Panophthalmitis; blindness Grade 5: death related to adverse event</p>										
<p>Adverse Event Heart- Late RT Morbidity Scoring</p> <p>Grade 1: Asymptomatic or mild symptoms; transient T wave inversion and ST changes; sinus tachycardia >110 (at rest) Grade 2: Moderate angina on effort; mild pericarditis; normal heart size; persistent abnormal T wave and ST changes; low QRS Grade 3: Severe angina; pericardial effusion; constrictive pericarditis; moderate heart failure; cardiac enlargement; EKG abnormalities Grade 4: Tamponade/severe heart failure/severe constrictive pericarditis Grade 5: death related to adverse event</p>										
<p>Adverse Event Joint- Late RT Morbidity Scoring</p> <p>Grade 1: Mild joint stiffness; slight limitation of movement Grade 2: Moderate stiffness; intermittent or moderate joint pain; moderate limitation of movement Grade 3: Severe joint stiffness; pain with severe limitation of movement Grade 4: Necrosis/complete fixation Grade 5: death related to adverse event</p>										
<p>Adverse Event Kidney-Late RT Morbidity Scoring</p> <p>Grade 1: Transient albuminuria; no hypertension; mild impairment of renal function; urea 25-35 mg%; creatinine 1.5-2.0 mg%; creatinine clearance >75% Grade 2: Persistent moderate albuminuria (2+); mild hypertension; no related anemia; moderate impairment of renal function; urea >36-60 mg%; creatinine clearance >50-74% Grade 3: Severe albuminuria; severe hypertension; persistent anemia (<10 g%); severe renal failure; urea >60 mg%; creatinine >4 mg%; creatinine clearance <50% Grade 4: Malignant hypertension; uremic coma/urea >100% Grade 5: death related to adverse event</p>										

in Morbidity Scoring Scheme (Use for adverse events occurring greater than 90 days after radiation therapy.)Category:	<p>Adverse Event Larynx-Late RT Morbidity Scoring</p> <p>Grade 1: Hoarseness; slight arytenoid edema Grade 2: Moderate arytenoid edema; chondritis Grade 3: Severe edema; severe chondritis Grade 4: Necrosis Grade 5: death related to adverse event</p>									
	<p>Adverse Event Liver-Late RT Morbidity Scoring</p> <p>Grade 1: Mild lassitude; nausea; dyspepsia; slightly abnormal liver function Grade 2: Moderate symptoms; some abnormal liver function tests; serum albumin normal Grade 3: Disabling hepatic insufficiency; liver function tests grossly abnormal; low albumin; edema or ascites Grade 4: Necrosis/hepatic coma or encephalopathy Grade 5: death related to adverse event</p>									
	<p>Adverse Event Lung-Late RT Morbidity Scoring</p> <p>Grade 1: Asymptomatic or mild symptoms (dry cough); slight radiographic appearances Grade 2: Moderate symptomatic fibrosis or pneumonitis (severe cough); low grade fever; patchy radiographic appearances Grade 3: Severe symptomatic fibrosis or pneumonitis; dense radiographic changes Grade 4: Severe respiratory insufficiency/continuous O(2)/assisted ventilation Grade 5: death related to adverse event</p>									
	<p>Adverse Event Mucous membrane-Late RT Morbidity Scoring</p> <p>Grade 1: Slight atrophy and dryness Grade 2: Moderate atrophy and telangiectasia; little mucus Grade 3: Marked atrophy with complete dryness; severe telangiectasia Grade 4: Ulceration Grade 5: death related to adverse event</p>									
	<p>Adverse Event Salivary glands-Late RT Morbidity Scoring</p> <p>Grade 1: Slight dryness of mouth; good response on stimulation Grade 2: Moderate dryness of mouth; poor response on stimulation Grade 3: Complete dryness of mouth; no response on stimulation Grade 4: Fibrosis Grade 5: death related to adverse event</p>									

Category: Appendix IV RTOG/EORTC Late Radiation	<p>Adverse Event Skin-Late RT Morbidity Scoring</p> <p>Grade 1: Slight atrophy; pigmentation change; some hair loss Grade 2: Patchy atrophy; moderate telangiectasia; total hair loss Grade 3: Marked atrophy; gross telangiectasia Grade 4: Ulceration Grade 5: death related to adverse event</p>									
	<p>Adverse Event Small/Large intestine-Late RT Morbidity Scoring</p> <p>Grade 1: Mild diarrhea; mild cramping; bowel movement 5 x daily; slight rectal discharge or bleeding Grade 2: Moderate diarrhea and colic; bowel movement >5 x daily; excessive rectal mucus or intermittent bleeding Grade 3: Obstruction or bleeding, requiring surgery Grade 4: Necrosis/perforation; fistula Grade 5: death related to adverse event</p>									
	<p>Adverse Event Spinal cord-Late RT Morbidity Scoring</p> <p>Grade 1: Mild Lhermitte's syndrome Grade 2: Severe Lhermitte's syndrome Grade 3: Objective neurological findings at or below cord level treatment Grade 4: Mono-, para-, quadriplegia Grade 5: death related to adverse event</p>									
	<p>Adverse Event Subcutaneous tissue-Late RT Morbidity Scoring</p> <p>Grade 1: Slight induration (fibrosis) and loss of subcutaneous fat Grade 2: Moderate fibrosis but asymptomatic; slight field contracture; <10% linear reduction Grade 3: Severe induration and loss of subcutaneous tissue; field contracture >10% linear measurement Grade 4: Necrosis Grade 5: death related to adverse event</p>									
	<p>Adverse Event Radiation-Other(Specify,_____)</p> <p>Grade 1: Mild Grade 2: Moderate Grade 3: Severe Grade 4: Life-threatening or disabling Grade 5: death related to adverse event</p>									
Appendix VI BMT Complex/Multicomponent Events										

	Toxicity category and code name	Description (reported term)	Grade	LLT MedDRA	Preferred term MedDRA	HLT	HLGT	Pr = Primary SOC Sec _n = Secondary SOC	SSC1	SSC2	SSC3
	<p>Note: The grading of Complex/Multicomponent Events in bone marrow transplant will be defined in the protocol. The grading scale must use the CTC criteria for grading the specific component events (toxicities). Category: Appendix VI BMT Complex/Multicomponent Events</p>										
<p>plex/Multicomponent Events</p>	<p><u>Adverse Event</u> Failure to engraft</p> <p><u>Grade 1</u>: mild <u>Grade 2</u>: moderate <u>Grade 3</u>: severe <u>Grade 4</u>: life-threatening <u>Grade 5</u>: death related to adverse event</p> <p>Note: Also consider Hemoglobin, Neutrophils/granulocytes (ANC/AGC), Neutrophils/granulocytes (ANC/AGC) for BMT studies, if specified in the protocol, Platelets, Platelets for BMT studies, if specified in the protocol.</p>										
	<p><u>Adverse Event</u> Graft versus host disease</p> <p><u>Grade 1</u>: mild <u>Grade 2</u>: moderate <u>Grade 3</u>: severe <u>Grade 4</u>: life-threatening <u>Grade 5</u>: death related to adverse event</p> <p>Note: Also consider Fatigue, Rash/desquamation, Rash/desquamation associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol, Diarrhea patients without a colostomy, Diarrhea patients with a colostomy, Diarrhea associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol, Diarrhea for pediatric BMT studies, if specified in the protocol, Bilirubin, Bilirubin associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol.</p>	<p>Graft versus host disease</p>	<p>1, 2, 3 or 4</p>	<p><u>LLT</u>: Graft versus host disease</p>	<p><u>PT</u>: Graft versus host disease</p>	<p><u>HLT</u>: Transplant rejections</p>	<p><u>HLGT</u>: Immune disorders NEC</p>	<p><u>Pr</u>: Immune system disorders</p>	<p><u>SSC</u>: SECONDARY IMMUNOCOM PROMISED STATE</p>		
<p><u>HLT</u>: Non-site specific procedural complications</p>	<p><u>HLGT</u>: Procedural and device related injuries and complications NEC</p>	<p><u>Sec₁</u>: Injury, poisoning and procedural complications</p>									

Category: Appendix VI BMT Comi	<p>Adverse Event Stem cell infusion complications</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening Grade 5: death related to adverse event</p> <p>Note: Also consider Allergic reaction/ hypersensitivity, Conduction abnormality/Atrioventricular heart block, Noda/junctional arrhythmia/dysrhythmia, Prolonged QTc interval (QTc >0.48 seconds), Sinus bradycardia, Sinus tachycardia, Supraventricular arrhythmias (SVT/atrial fibrillation/flutter), Vasovagal episode, Ventricular arrhythmia (PVCs/bigeminy/trigeminy/ventricular tachycardia), Cardiovascular/Arrhythmia-Other (Specify, _____), Hypertension, Hypotension, Fever (in the absence of neutropenia, where neutropenia is defined as AGC <1.0 x 10(9)/L), Rigors/chills, Sweating (diaphoresis), Rash/desquamation, Rash/desquamation associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol, Urticaria (hives, welts, wheals), Diarrhea patients without a colostomy, Diarrhea patients with a colostomy, Diarrhea associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol</p>									
	<p>Adverse Event Veno-Occlusive Disease (VOD)</p> <p>Grade 1: mild Grade 2: moderate Grade 3: severe Grade 4: life-threatening Grade 5: death related to adverse event</p> <p>Note: Also consider Weight gain associated with Veno-Occlusive Disease (VOD) for BMT studies, if specified in the protocol, Bilirubin, Bilirubin associated with graft versus host disease (GVHD) for BMT studies, if specified in the protocol, Depressed level of consciousness, Hepatic pain, Renal failure, Hepatic enlargement.</p>	Veno-Occlusive Disease (VOD)	1, 2, 3 or 4	LLT: Venoocclusive disease NOS	PT: Venoocclusive disease NOS	HLT: Non-site specific vascular disorders NEC	HLGT: Vascular disorders NEC	Pr: Vascular disorders		