B120719

Day numbers relative to Start Date

					-16	-15	-14	-13	-12	-11	-10	- 9	- 8
Group		Animal	Clinical Sign	Site	a.m.	a.m.							
												• • • • • • • • • • • • • • • • • • • •	
2	m	10201	No Abnormalities Detected		X	Х	Х	Х	Х	Х	Х	Х	Х
		10202	No Abnormalities Detected		Х	Х	Х	Х	Х	X	X	Х	Х
		10203	No Abnormalities Detected		Х	X	Х	Х	Х	Х	X	X	Х

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Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

B120719

Day numbers relative to Start Date

					- 7	-6	- 5	- 4	-3	- 2	- 1
Group	Sex	Animal	Clinical Sign Site	а	. m.	a.m.	a.m.	a.m.	a.m.	a.m.	a.m.
2	m	10201	No Abnormalities Detected		Х	Х	X	Х	Х	Х	X
		10202	No Abnormalities Detected		Χ	Х	Х	X	Х	X	X
		10203	No Abnormalities Detected		Х	X	Х	Х	X	X	X

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg Time 10 , 2h after dosing;

B120719

Day numbers relative to Start Date

Grou	p Sex	Animal	Clinical Sign	Site	-52 a.m.	-51 a.m.	-50 a.m.	-49 a.m.	-48 a.m.	-47 a.m.	-46 a.m.	-45 a.m.	-44 a.m.
3	m	10301 10302	No Abnormalities Detected No Abnormalities Detected		X X								
		10303	No Abnormalities Detected		X	X	x	Х	X	X	X	X	X

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

B120719

Day numbers relative to Start Date

					- 43	- 42	- 41	-40	- 39	- 38	-37	-36	-35
Grou	p Sex	Animal	Clinical Sign	Site	a.m.	a.m.	a.m.	a.m.	a.m.	a.m.	a.m.	a.m.	a.m.
										. 			
3	m	10301	No Abnormalities Detected		X	Х	X	X	X	Х	X	X	Х
		10302	No Abnormalities Detected		Х	Х	X	X	Х	Х	X	X	Х
		10303	No Abnormalities Detected		Х	X	X	X	Х	X	X	X	Х

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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B120719

Day numbers relative to Start Date

					-34	- 33	-32	- 31	-30	- 29	-28	- 27	- 26
	ıp Sex	Animal	Clinical Sign	Site	a.m.								
3	m	10301	No Abnormalities Detected		χ	χ	Х	Х	Х	χ	Х	Х	χ
		10302	No Abnormalities Detected		X	Х	Х	X	X	Х	X	X	X
		10303	No Abnormalities Detected		X	Х	X	Х	X	X	X	X	X

.....

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

B120719

Day numbers relative to Start Date

Group		Animal	Clinical Sign	Site	-25 a.m.	-24 a.m.	-23 a.m.	-22 a.m.	-21 a.m.	-20 a.m.	-19 a.m.	-18 a.m.	-17 a.m.
3	m	10301 10302 10303	No Abnormalities Detected No Abnormalities Detected No Abnormalities Detected		X X X								

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

					-16	-15	-14	-13	-12	-11	-10	- 9	-8
Group		Animal	Clinical Sign	Site	a.m.								
3	m	10301	No Abnormalities Detected		Х	Х	Х	Х	х	Х	Х	Х	X
		10302	No Abnormalities Detected		Х	Х	Х	Х	X	Х	X	Х	X
		10303	No Abnormalities Detected		X	Х	Х	Х	Х	X	X	Х	X

.....

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

B120719

Day numbers relative to Start Date

					-7	-6	- 5	- 4	-3	-2	- 1
Gro	up Sex	Animal	Clinical Sign	Site	a.m.						
3	m	10301	No Abnormalities Detected		х	Х	Х	Х	Х	X	X
		10302	No Abnormalities Detected		Х	Х	Х	Х	Х	Х	Х
		10303	No Abnormalities Detected		X	Х	Х	X	Х	X	X

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg Time 10, 2h after dosing;

B120719

Day numbers relative to Start Date

					-52	- 51	-50	- 49	- 48	- 47	- 46	- 45	-44
Grou	p Sex	Animal	Clinical Sign	Site	a.m.	a.m.							
4	m	10401	No Abnormalities Detected		X	X	X	X	X	Х	X	. X	Х
		10402	No Abnormalities Detected		X	X	X	Х	X	Х	X	X	X
		10403	No Abnormalities Detected		X	Χ	X	X	X	X	X	X	Х

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

	•	Animal	Clinical Sign	Site	-43 a.m.	-42 a.m.	-41 a.m.	-40 a.m.	-39 a.m.	-38 a.m.	-37 a.m.	-36 a.m.	-35 a.m.
4	m	10401 10402 10403	No Abnormalities Detected No Abnormalities Detected No Abnormalities Detected		X X X								

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

	•	Animal	Clinical Sign	Site	-34 a.m.	-33 a.m.	-32 a.m.	-31 a.m.	-30 a.m.	-29 a.m.	-28 a.m.	-27 a.m.	-26 a.m.
4	m	10401 10402 10403	No Abnormalities Detected No Abnormalities Detected No Abnormalities Detected		X X X								

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

Appendix 1

Clinical Sign

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Day numbers relative to Start Date

Group Sex Animal	Clinical Sign	Site	-25 a.m.	-24 a.m.	-23 a.m.	-22 a.m.	-21 a.m.	-20 a.m.	-19 a.m.	-18 a.m.	-17 a.m.
4 m 10401	No Abnormalities Detected		X	X	X	X	X	X	X	X	X
10402	No Abnormalities Detected		X	X	X	X	X	X	X	X	X
10403	No Abnormalities Detected		Y	Y	Y	Y	Y	Y	Y	Y	X

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

[Dose level was changed from 500 mg/kg to 250 mg/kg on Day 5.]

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Day numbers relative to Start Date

					-16	-15	- 14	-13	-12	-11	-10	-9	-8
Group		Animal	Clinical Sign	Site	a.m.								
4	 m	10401	No Abnormalities Detected		Х	Х	X	Х	Х	Х	X	X	χ
		10402	No Abnormalities Detected		X	Х	Х	Х	X	X	Х	Х	X
		10403	No Abnormalities Detected		X	X	X	X	X	Х	X	X	X

.....

Severity Codes: X = Present

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

				-7	-6	-5	- 4	-3	- 2	- 1
Group	p Sex	Animal	Clinical Sign Site	a.m.	a.m.	a.m.	a.m.	a.m.	a.m.	a.m.
4	m	10401	No Abnormalities Detected	Х	X	Х	Х	Х	Х	Х
		10402	No Abnormalities Detected	. X	Х	Х	Х	X	X	X
		10403	No Abnormalities Detected	Х	Х	Х	Х	Х	Х	Х

Severity Codes: X = Present

Time 10, 2h after dosing;

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Day numbers relative to Start Date

					1	1	2	2	3	3	4	4	5
Group :		Animal	Clinical Sign	Site	Before dosing	Time 10	Before dosing						
1	m	10101	No Abnormalities Detected		X	X	Х	Х	Х	X	Х	X	χ
		10102	No Abnormalities Detected		X	X	Х	Х	X	Х	Х	X	X
		10103	No Abnormalities Detected		Х	Х	Х	X	Х	Х	Х	Х	Х

Severity Codes: X = Present; 1 = Slight

Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg Group 1 - 0 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

Group Sex Animal	Clinical Sign Site	5 Time 10	6 Before dosing	6 Time 10	7 Before dosing	7 Time 10	8 Before dosing	8 Time 10	9 Before dosing	9 Time 10
1 m 10101	No Abnormalities Detected	х	X	X	X	X	X	X	X	X
10102	No Abnormalities Detected	х	X	X	X	X	X	X	X	X
10103	No Abnormalities Detected	х	X	X	X	X	X	X	X	X

Severity Codes: X = Present; 1 = Slight

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

					10	10	11	11	12	12	13	13	14
Group		Animal	Clinical Sign	Site	Before dosing	Time 10	Before dosing						
1	m		No Abnormalities Detected		Х	X	х	Х	Х	X	Х	Х	X
		10102	No Abnormalities Detected		Х	X	X	X	Х	Х	Х	X	X
		10103	No Abnormalities Detected		Х	Х	Х	Х	Х	Х	Х	X	Х

Severity Codes: X = Present; 1 = Slight

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

						14	15
						Time 10	a.m.
Group	Sex	Animal	Clinical	Sign	Site		
1	 m	10101	No Abnormalities	Detected		X	Х
		10102	No Abnormalities	Detected		X	Х
		10103	No Abnormalities	Detected		X	X

Severity Codes: X = Present; 1 = Slight

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

[Dose level was changed from 500 mg/kg to 250 mg/kg on Day 5.]

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Day numbers relative to Start Date

Group Sex Animal	Clinical Sign	Site	1 Before dosing	1 Time 10	2 Before dosing	2 Time 10	3 Before dosing	3 Time 10	4 Before dosing	4 Time 10	5 Before
Group Sex Annual	Central Sign	31te	aosing		ausing		ausing		aosing		dosing
2 m 10201	No Abnormalities Detected		Х	X					Х		X
	Vomitus (Dosing formulation)									1	
	Vomitus (Food)		•		Х		Х				
	Vomitus (Frothy material)			•				-		•	
	Vomitus (Dosing solution)					Х	-	Х		•	•
10202	No Abnormalities Detected		Х			Х		•	X		X
	Vomitus (Dosing formulation)			•			•			1	•
	Vomitus (Food)			•	Х		Х				•
	Vomitus (Dosing solution)			Х	-					•	
	Diarrhea			•				Х	-		
10203	No Abnormalities Detected		Х		Х		Х	•	Х	•	X
	Vomitus (Dosing formulation)			•		-				1	
	Vomitus (Food)					-	-				
	Vomitus (Frothy material)				•	-		•		•	
	Vomitus (Dosing solution)			Х		Х	•	Х		•	

Severity Codes: X = Present; 1 = Slight

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;

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Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	5 Time 10	6 Before dosing	6 Time 10	7 Before dosing	7 Time 10	8 Before dosing	8 Time 10	9 Before dosing	9 Time 10
2	m	10201	No Abnormalities Detected			χ		Х		X		Х	X
			Vomitus (Dosing formulation)				1			•		-	
			Vomitus (Food)		•			•		•	•	•	
			Vomitus (Frothy material)								Х	-	
			Vomitus (Dosing solution)		Х				X	-	. •		
		10202	No Abnormalities Detected			Х		Х		Х		Х	Х
			Vomitus (Dosing formulation)				1	ь		•	•		
•			Vomitus (Food)					•				-	
			Vomitus (Dosing solution)		X				Х		X	-	
			Diarrhea				•	-		•		-	
		10203	No Abnormalities Detected			Х		Х		X		X	
			Vomitus (Dosing formulation)				1					-	-
			Vomitus (Food)			•	•					-	
			Vomitus (Frothy material)			•		•					Х
			Vomitus (Dosing solution)		Х		•		Х		Х		

Severity Codes: X = Present; 1 = Slight

Group 1 - 0 mg/kg Group 2 - 50 mg/kg Group 3 - 150 mg/kg Group 4 - 500 mg/kg

Time 10, 2h after dosing;