

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 3 Food consumption (males)

Sex: Male Food consumption (g/day)

Low-Mid TM5509 6 mg/kg/day	Day(s) Relative to Start Date													
	15	16	17	18	19	20	21	22	23	24	25	26	27	28
RP3M01	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP3M02	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP3M03	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP3M04	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 3 Food consumption (males)

Sex: Male Food consumption (g/day)

Mid TM5509 20 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP4M01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP4M02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP4M03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP4M04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 3 Food consumption (males)

Sex: Male		Food consumption (g/day)													
Mid TM5509 20 mg/kg/day	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP4M01	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP4M02	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP4M03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP4M04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4	

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Appendix 3 Food consumption (males)

Sex: Male		Food consumption (g/day)																	
Mid-High TM5509 80 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP5M01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP5M02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP5M03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP5M04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 3 Food consumption (males)

Sex: Male		Food consumption (g/day)													
Mid-High TM5509 80 mg/kg/day	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP5M01	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP5M02	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP5M03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP5M04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4	

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Appendix 3 Food consumption (males)

Sex: Male Food consumption (g/day)

High TM5509 300 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP6M01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP6M02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	80
RP6M03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP6M04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	95
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	10
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 3 Food consumption (males)

Sex: Male		Food consumption (g/day)													
High TM5509 300 mg/kg/day	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP6M01	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP6M02	0	
RP6M03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP6M04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	75	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	50	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	4	3	3	3	3	3	3	3	3	3	3	3	3	3	

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)																		
Control 0.5%CMC-Na	Day(s) Relative to Start Date																			
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14	
RP1F01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP1F02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP1F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP1F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	

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A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)													
Control 0.5%CMC-Na	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP1F01	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP1F02	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP1F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP1F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4	

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Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)																	
Low TM5509 2 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP2F01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP2F02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP2F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP2F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 4 Food consumption (females)

Sex: Female

Food consumption (g/day)

Low TM5509 2 mg/kg/day	Day(s) Relative to Start Date													
	15	16	17	18	19	20	21	22	23	24	25	26	27	28
RP2F01	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP2F02	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP2F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP2F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 4 Food consumption (females)

Sex: Female Food consumption (g/day)

Low-Mid TM5509 6 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP3F01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP3F02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP3F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP3F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)													
Low-Mid TM5509 6 mg/kg/day	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP3F01	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP3F02	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP3F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP3F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4	

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 4 Food consumption (females)

Sex: Female

Food consumption (g/day)

Mid TM5509 20 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP4F01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP4F02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP4F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP4F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)													
Mid TM5509 20 mg/kg/day	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP4F01	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP4F02	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP4F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP4F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4	

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)																	
Mid-High TM5509 80 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP5F01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP5F02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP5F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP5F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)													
Mid-High TM5509 80 mg/kg/day	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP5F01	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP5F02	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP5F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP5F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
N	4	4	4	4	4	4	4	4	4	4	4	4	4	4	

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 4 Food consumption (females)

Sex: Female

Food consumption (g/day)

High TM5509 300 mg/kg/day	Day(s) Relative to Start Date																		
	-5	-4	-3	-2	-1	1	2	3	4	5	6	7	8	9	10	11	12	13	14
RP6F01	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP6F02	.	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP6F03	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
RP6F04	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Mean	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
S.D.	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
N	2	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4

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Appendix 4 Food consumption (females)

Sex: Female		Food consumption (g/day)													
High TM5509 300 mg/kg/day	Day(s) Relative to Start Date														
	15	16	17	18	19	20	21	22	23	24	25	26	27	28	
RP6F01	100	100	100	100	0	
RP6F02	100	100	100	100	100	100	100	100	100	100	100	100	100	100	
RP6F03	100	100	100	100	
RP6F04	100	100	100	100	
Mean	100	100	100	100	50	100	100	100	100	100	100	100	100	100	
S.D.	0	0	0	0	71	
N	4	4	4	4	2	1	1	1	1	1	1	1	1	1	

A 4-Week Oral Dose Toxicity Study of TM5509 in Cynomolgus Monkeys

Appendix 5 Hematology (males)

Sex: Male Week(s) Relative to Start Date

Control 0.5%CMC-Na	RBC					HGB				
	(10 ⁶ /μL)	(10 ⁶ /μL)	(10 ⁶ /μL)	(10 ⁶ /μL)	(10 ⁶ /μL)	(g/dL)	(g/dL)	(g/dL)	(g/dL)	(g/dL)
	-1	1	2	3	4	-1	1	2	3	4
RP1M01	5.35	5.26	5.27	5.24	5.28	13.2	12.8	12.9	12.9	12.8
RP1M02	6.24	5.69	5.74	5.73	5.91	15.0	13.7	13.9	13.9	14.2
RP1M03	5.21	4.87	4.79	4.86	4.79	14.1	13.1	13.0	13.1	12.9
RP1M04	6.65	6.48	6.50	6.22	6.30	15.7	15.2	15.3	14.6	14.7
Mean	5.86	5.58	5.58	5.51	5.57	14.5	13.7	13.8	13.6	13.7
S.D.	0.70	0.69	0.73	0.59	0.67	1.1	1.1	1.1	0.8	0.9
N	4	4	4	4	4	4	4	4	4	4