

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

Table 1 Clinical observations (males)

Sex: Male

Group	Animal No.	Findings	Day of dosing																NE am								
			20			21			22			23			24			25			26			27			NE am
			Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	
Control 0.5%CMC-Na	RP1M01	No abnormalities																									
	RP1M02	No abnormalities																									
	RP1M03	No abnormalities																									
	RP1M04	No abnormalities																									
Low TM5509 2 mg/kg/day	RP2M01	No abnormalities																									
	RP2M02	No abnormalities																									
	RP2M03	No abnormalities																									
	RP2M04	No abnormalities																									
Low-Mid TM5509 6 mg/kg/day	RP3M01	No abnormalities																									
	RP3M02	No abnormalities																									
	RP3M03	No abnormalities																									
	RP3M04	No abnormalities																									
Mid TM5509 20 mg/kg/day	RP4M01	No abnormalities																									
	RP4M02	No abnormalities																									
	RP4M03	No abnormalities																									
	RP4M04	No abnormalities																									
Mid-High TM5509 80 mg/kg/day	RP5M01	No abnormalities																									
	RP5M02	Muddy stools	(normal feces: present)																								
	RP5M03	Muddy stools	(normal feces: absent)																								
	RP5M04	Muddy stools	(normal feces: present)																								
High TM5509 300 mg/kg/day	RP6M01	Grayish white muddy stools	(normal feces: absent)																								
			(normal feces: present)																								
		Vomitus (test article-like substances):	Moderate quantity																		A						
		Grayish white stools	(normal feces: present)																								
		Muddy stools	(normal feces: present)																			A					

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing NE: Day of necropsy A: Appearance of signs Blank spaces indicate "no abnormalities";

a) 4 and 6 hours post-dosing b) 8 hours post-dosing

a) 4 and 6 hours post-dosing b) 8 hours post-dosing

NE: Day of necropsy

A: Appearance of signs

Blank spaces indicate "no abnormalities".

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

Table 1 Clinical observations (males)

Sex: Male

Group	Animal No.	Findings	Day of dosing												NE															
			20			21			22			23			24			25			26			27			28			
			Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	am			
High TM5509 300 mg/kg/day	RP6M02	Vomiting (test article-like substances): Small quantity																												
		Large quantity																												
		Grayish white muddy stools (normal feces: absent)																												
		Grayish white loose stools (normal feces: absent)																												
		(normal feces: present)																												
		Grayish white stools (normal feces: present)																												
		Muddy stools (normal feces: absent)																												
		(normal feces: present)																												
		Decrease in movement																												
		Vomitus (undigested feed)																												
		(test article-like substances)																												
		Found dead																												
RP6M03	RP6M03	Grayish white watery stools (normal feces: absent)																												
		Lateral position																												
		Grayish white loose stools (normal feces: present)	A			A			A							A		A		A				A						
		Grayish white stools (normal feces: present)																												
		Grayish white muddy stools (normal feces: absent)																						A						
		(normal feces: present)																												
		Muddy stools (normal feces: absent)																							A ^{a)}					
		(normal feces: present)																												
		No feces																												
		Vomitus (undigested feed)																												
		Grayish white muddy stools (normal feces: absent)																												
		(normal feces: present)																												
RP6M04	RP6M04	Muddy stools (normal feces: present)																												
		Grayish white stools (normal feces: absent)																												
		(normal feces: present)																												
		Grayish white stools (normal feces: present)																							A ^{a)}					

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing NE: Day of necropsy A: Appearance of signs Blank spaces indicate "no abnormalities".

^{a)} 8 hours post-dosing

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

Table 2 Clinical observations (females)

Sex: Female

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs Blank spaces indicate "no abnormalities".

a) Immediately after dosing

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

Table 2 Clinical observations (females)

Sex: Female

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs Blank spaces indicate "no abnormalities".

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

Table 2 Clinical observations (females)

Sex: Female

Group	Animal No.	Findings	Day of dosing																													
			-6or-5to-1			1			2			3			4			5			6			7			8			9		
			am	Pre	1h	4h																										
High TM5509 300 mg/kg/day	RP6F04	Grayish white muddy stools (normal feces: present)											A			A																
		(normal feces: absent)																														
		Loose stools (normal feces: present)																														
		Muddy stools (normal feces: absent)																														
		(normal feces: present)																														
		Menses																														
		Vomitus (test article-like substances)																														
		(undigested feed)																														
		Vomiting (test article-like substances): Moderate quantity																														
		Grayish white stools (normal feces: present)																														
		Subnormal body surface temperature																														
		Watery stools (normal feces: absent)																														
		Moribund condition																														
		Residual feed																														

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs Blank spaces indicate "no abnormalities".

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

Table 2 Clinical observations (females)

Sex: Female

Group	Animal No.	Findings	Day of dosing																	
			10		11		12		13		14		15		16		17		18	
Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h
Control	RP1F01	Menses																		
0.5%CMC-Na	RP1F02	No abnormalities																		
	RP1F03	No abnormalities																		
	RP1F04	Menses																		
Low	RP2F01	No abnormalities																		
TM5509	RP2F02	Menses																		
2	RP2F03	No abnormalities																		
mg/kg/day	RP2F04	Menses																		
Low-Mid	RP3F01	No abnormalities																		
TM5509	RP3F02	No abnormalities																		
6	RP3F03	No abnormalities																		
mg/kg/day	RP3F04	No abnormalities																		
Mid	RP4F01	Menses																		
TM5509	RP4F02	Menses																		
20	RP4F03	No abnormalities																		
mg/kg/day	RP4F04	Menses							A	A	A	A	A	A	A	A	A	A	A	
Mid-High	RP5F01	Vomiting (test article-like substances): Small quantity																		
TM5509		Menses	A	A	A	A	A	A	A	A	A	A	A	A	A	A	A	A	A	
80	RP5F02	Muddy stools (normal feces: present)																		
mg/kg/day		(normal feces: absent)																		
	RP5F03	Muddy stools (normal feces: present)																		
	RP5F04	Muddy stools (normal feces: present)																		

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs Blank spaces indicate "no abnormalities".

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

Table 2 Clinical observations (females)

Sex: Female

Group	Animal No.	Findings	Day of dosing																			
			10				11				12				13				14			
			Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h		
High TM5509 300 mg/kg/day	RP6F01	Grayish white muddy stools (normal feces: present)													A			A				
		(normal feces: absent)																				
		Grayish white loose stools (normal feces: present)													A							
		Grayish white stools (normal feces: present)																				
		Muddy stools (normal feces: absent)													A	A						
		Menses																				
		Vomitus (test article-like substances)													A							
		Vomiting (test article-like substances): Moderate quantity																	A			
		No feces																				
		Subnormal body surface temperature																				
		Prone position																				
		Moribund condition																				
RP6F02		Grayish white loose stools (normal feces: present)																				
		Grayish white stools (normal feces: present)													A							
		Grayish white muddy stools (normal feces: absent)													A							
		(normal feces: present)																		A		
RP6F03		Grayish white muddy stools (normal feces: present)														A	A	A	A	A		
		(normal feces: absent)													A							
		Vomitus (undigested feed)	A																	A		
		Grayish white stools (normal feces: present)													A	A				A		
		Vomitus (test article-like substances)													A					A		
		Vomiting (test article-like substances): Moderate quantity																A ^{b)}				
		Subnormal body surface temperature																		A ^{c)}		
		Moribund condition ^{a)}																		A		
		Subnormal body surface temperature																		A ^{d)}		
		Light yellow discoloration of the oral mucosa																		A		
		Residual feed																		A ^{e)}		

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs Blank spaces indicate "no abnormalities".
 a) Necropsied b) Immediately after dosing c) Body temperature: 36.7 °C d) Body temperature: 36.2 °C e) 30 g

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

Table 2 Clinical observations (females)

Sex: Female

General Footnote: Pre; Pre-dosing 1h and 4h; 1 and 4 hours post-dosing A; Appearance of signs Blank spaces indicate "no abnormalities".

a) Necropsied

1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs

Blank spaces indicate "no abnormalities".

a) Necropsied

1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs

Blank spaces indicate "no abnormalities".

a) Necropsied

1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs

Blank spaces indicate "no abnormalities".

a) Necropsied

1h and 4h: 1 and 4 hours post-dosing A: Appearance of signs

Blank spaces indicate "no abnormalities".

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

Table 2 Clinical observations (females)

Sex: Female

Group	Animal No.	Findings	Day of dosing																NE							
			20			21			22			23			24			25			26			9		
			Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	am		
Control	RP1F01	Menses																								
0.5%CMC-Na	RP1F02	No abnormalities																								
	RP1F03	No abnormalities																								
	RP1F04	Menses																								
Low	RP2F01	No abnormalities																								
TM5509	RP2F02	Menses																								
2	RP2F03	No abnormalities																								
mg/kg/day	RP2F04	Menses																					A	A	A	A
Low-Mid	RP3F01	No abnormalities																								
TM5509	RP3F02	No abnormalities																								
6	RP3F03	No abnormalities																								
mg/kg/day	RP3F04	No abnormalities																								
Mid	RP4F01	Menses																				A	A	A	A	
TM5509	RP4F02	Menses																				A	A	A	A	
20	RP4F03	No abnormalities																				A	A	A	A	
mg/kg/day	RP4F04	Menses																				A	A	A	A	
Mid-High	RP5F01	Vomiting (test article-like substances): Small quantity																								
TM5509		Menses																								
80	RP5F02	Muddy stools (normal feces: present)																								
mg/kg/day		(normal feces: absent)																								
	RP5F03	Muddy stools (normal feces: present)																								
	RP5F04	Muddy stools (normal feces: present)																								

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing NE: Day of necropsy A: Appearance of signs Blank spaces indicate "no abnormalities".

^{a)} 8 hours post-dosing

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

Table 2 Clinical observations (females)

Sex: Female

Group	Animal No.	Findings	Day of dosing												NE									
			20	21	22	23	24	25	26	27	28	Pre	1h	4h	am									
High TM5509 300 mg/kg/day	RP6F01	Grayish white muddy stools (normal feces: present)																						
		(normal feces: absent)																						
		Grayish white loose stools (normal feces: present)																						
		Grayish white stools (normal feces: present)																						
		Muddy stools (normal feces: absent)																						
		Menses																						
		Vomitus (test article-like substances)																						
		Vomiting (test article-like substances): Moderate quantity																						
		No feces	A																					
		Subnormal body surface temperature	A ^{b)}																					
		Prone position	A																					
		Moribund condition ^{a)}	A																					
RP6F02		Grayish white loose stools (normal feces: present)																						
		Grayish white stools (normal feces: present)																						
		Grayish white muddy stools (normal feces: absent)																						
		(normal feces: present)																						
RP6F03	RP6F03	Grayish white muddy stools (normal feces: present)																						
		(normal feces: absent)																						
		Vomitus (undigested feed)																						
		Grayish white stools (normal feces: present)																						
		Vomitus (test article-like substances)																						
		Vomiting (test article-like substances): Moderate quantity																						
		Subnormal body surface temperature																						
		Moribund condition																						
		Subnormal body surface temperature																						
		Light yellow discoloration of the oral mucosa																						
		Residual feed																						

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing NE: Day of necropsy A: Appearance of signs Blank spaces indicate "no abnormalities".

^{a)} This animal was determined to be killed, but died prior to removal from the animal room. ^{b)} Body temperature: 34.1 °C

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

Table 2 Clinical observations (females)

Sex: Female

Group	Animal No.	Findings	Day of dosing												NE				
			20		21		22		23		24		25		26		27		
			Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	Pre	1h	4h	am	
High TM5509 300 mg/kg/day	RP6F04	Grayish white muddy stools	(normal feces: present)																
			(normal feces: absent)																
		Loose stools	(normal feces: present)																
		Muddy stools	(normal feces: absent)																
			(normal feces: present)																
		Menses																	
		Vomitus	(test article-like substances)																
			(undigested feed)																
		Vomiting	(test article-like substances): Moderate quantity																
		Grayish white stools	(normal feces: present)																
		Subnormal body surface temperature																	
		Watery stools	(normal feces: absent)																
		Moribund condition																	
		Residual feed																	

General Footnote: Pre: Pre-dosing 1h and 4h: 1 and 4 hours post-dosing NE: Day of necropsy

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

Table 3 Body weight (males)

Body weight (kg)

Sex: Male		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
1	Mean	3.24 ¹⁾	3.14	3.22	3.16	3.03	3.11
	S.D.	0.30	0.33	0.42	0.44	0.35	0.52
	N	4	4	4	4	4	4
4	Mean	3.26 ¹⁾	3.17	3.29	3.20	3.06	3.04
	S.D.	0.32	0.34	0.40	0.45	0.32	0.47
	N	4	4	4	4	4	4
8	Mean	3.25 ¹⁾	3.16	3.28	3.22	3.07	3.05
	S.D.	0.33	0.37	0.42	0.47	0.37	0.51
	N	4	4	4	4	4	4
11 or 12	Mean	3.31 ¹⁾	3.19	3.32	3.31	3.13	3.01
	S.D.	0.33	0.37	0.44	0.49	0.40	0.46
	N	4	4	4	4	4	4
15	Mean	3.30 ¹⁾	3.19	3.26	3.25	3.07	2.89
	S.D.	0.34	0.36	0.43	0.45	0.40	0.37
	N	4	4	4	4	4	4
18	Mean	3.35 ¹⁾	3.19	3.26	3.21	3.12	2.84
	S.D.	0.34	0.35	0.43	0.47	0.42	0.33
	N	4	4	4	4	4	3

Statistical Test: Generalised Anova/Ancova Test Transformation: Automatic

1 [1 - Automatic Transformation Selected: Identity (No Transformation)]

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

Table 3 Body weight (males)

Body weight (kg)

Sex: Male		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
22	Mean	3.38 ¹⁾	3.20	3.28	3.27	3.11	2.82
	S.D.	0.32	0.39	0.41	0.48	0.44	0.36
	N	4	4	4	4	4	3
25	Mean	3.37 ¹⁾	3.21	3.29	3.28	3.11	2.82
	S.D.	0.34	0.38	0.46	0.48	0.43	0.36
	N	4	4	4	4	4	3
28	Mean	3.42 ¹⁾	3.22	3.44	3.31	3.19	2.88
	S.D.	0.31	0.38	0.43	0.49	0.41	0.33
	N	4	4	4	4	4	3

Statistical Test: Generalised Anova/Ancova Test Transformation: Automatic

1 [1 - Automatic Transformation Selected: Identity (No Transformation)]

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

Table 4 Body weight (females)

Body weight (kg)

Sex: Female		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
1	Mean	2.73 ¹⁾	2.72	2.76	2.66	2.71	2.69
	S.D.	0.35	0.31	0.26	0.33	0.38	0.26
	N	4	4	4	4	4	4
4	Mean	2.74 ¹⁾	2.73	2.75	2.62	2.68	2.65
	S.D.	0.39	0.29	0.25	0.32	0.34	0.29
	N	4	4	4	4	4	4
8	Mean	2.78 ¹⁾	2.77	2.72	2.62	2.69	2.55
	S.D.	0.35	0.32	0.22	0.30	0.31	0.29
	N	4	4	4	4	4	4
11	Mean	2.82 ¹⁾	2.80	2.74	2.70	2.70	2.55
	S.D.	0.39	0.33	0.25	0.29	0.33	0.29
	N	4	4	4	4	4	4
15	Mean	2.82 ¹⁾	2.80	2.72	2.64	2.63	2.49
	S.D.	0.39	0.30	0.28	0.28	0.35	0.31
	N	4	4	4	4	4	4
18	Mean	2.83 ¹⁾	2.82	2.70	2.66	2.68	2.41
	S.D.	0.43	0.35	0.27	0.30	0.33	0.33
	N	4	4	4	4	4	4

Statistical Test: Generalised Anova/Ancova Test Transformation: Automatic

1 [I - Automatic Transformation Selected: Identity (No Transformation)]

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

Table 4 Body weight (females)

Body weight (kg)

Sex: Female		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
22	Mean	2.84 1 ¹	2.85	2.74	2.68	2.72	2.65 n1 ²
	S.D.	0.39	0.37	0.28	0.28	0.36	.
	N	4	4	4	4	4	1
25	Mean	2.87 1 ¹	2.85	2.78	2.79	2.78	2.73 n1 ²
	S.D.	0.43	0.40	0.26	0.39	0.35	.
	N	4	4	4	4	4	1
28	Mean	2.91 1 ¹	2.87	2.80	2.79	2.86	2.69 n1 ²
	S.D.	0.39	0.41	0.28	0.37	0.37	.
	N	4	4	4	4	4	1

Statistical Test: Generalised Anova/Ancova Test Transformation: Automatic

1 [1 - Automatic Transformation Selected: Identity (No Transformation)]

2 [n1 - This group has only one value: Data was not statistically analyzed.]

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

Table 5 Food consumption (males)

Food consumption (g/day)

Sex: Male		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
-4	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
-3	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
-2	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
-1	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
1	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
2	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
3	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4

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

Table 5 Food consumption (males)

Food consumption (g/day)

Sex: Male		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
4	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
5	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
6	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
7	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
8	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
9	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
10	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4

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

Table 5 Food consumption (males)

Food consumption (g/day)

Sex: Male		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
11	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
12	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
13	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	4
14	Mean	100 ¹	100	100	100	100	95
	S.D.	0	0	0	0	0	10
	N	4	4	4	4	4	4
15	Mean	100 ¹	100	100	100	100	75
	S.D.	0	0	0	0	0	50
	N	4	4	4	4	4	4
16	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
17	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3

Statistical Test: Generalised Anova/Ancova Test Transformation: Automatic

¹ [I - Automatic Transformation Selected: Identity (No Transformation)]

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

Table 5 Food consumption (males)

Food consumption (g/day)

Sex: Male		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
18	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
19	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
20	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
21	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
22	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
23	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
24	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3

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

Table 5 Food consumption (males)

Food consumption (g/day)

Sex: Male		Control 0.5%CMC-Na	Low TM5509 2 mg/kg/day	Low-Mid TM5509 6 mg/kg/day	Mid TM5509 20 mg/kg/day	Mid-High TM5509 80 mg/kg/day	High TM5509 300 mg/kg/day
Day(s) Relative to Start Date							
25	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
26	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
27	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3
28	Mean	100	100	100	100	100	100
	S.D.	0	0	0	0	0	0
	N	4	4	4	4	4	3