Test Type: 26-WEEK
Route: DOSED FEED

C Number:

Species/Strain: Mouse/P53(C57BL/6)

P11: STATISTICAL ANALYSIS OF SURVIVAL DATA

Test Compound: Transgenic model evaluation (Methylphenidate hydrochloride)

CAS Number: 298-59-9

First Dose M/F: NA / NA

Date Report Requested: 10/17/2014

Time Report Requested: 22:20:28

Lab: BAT

C95007V

Lock Date: 08/12/1996

Cage Range: All

Date Range: All

Reasons For Removal:

Removal Date Range: All

Treatment Groups: All

Study Gender: Both

PWG Approval Date NONE

Species/Strain: Mouse/P53(C57BL/6)

Test Type: 26-WEEK

Route: DOSED FEED

P11: STATISTICAL ANALYSIS OF SURVIVAL DATA

Test Compound: Transgenic model evaluation (Methylphenidate hydrochloride)

CAS Number: 298-59-9

Date Report Requested: 10/17/2014
Time Report Requested: 22:20:28

First Dose M/F: NA / NA

Lab: BAT

Male MOUSE FIRST TERMINAL SACRIFICE AT 169 DAYS INDIVIDUAL SURVIVAL TIMES (DAYS)

| | IIID | IVIDUAL SURVIVAL TIMES (DAT | 3) | |
|-----------------------|---------------------|-----------------------------|-------------|--|
| DOSE = 0 PPM | | | | |
| TOTAL 15 | UNCENSORED DEATHS 0 | CENSORED DEATHS 0 | TERMINAL 15 | |
| UNCENSORED DEATH DAYS | | | | |
| none | | | | |
| CENSORED DEATH DAYS | | | | |
| none | | | | |
| DOSE = 50 PPM | | | | |
| TOTAL 15 | UNCENSORED DEATHS 0 | CENSORED DEATHS 0 | TERMINAL 15 | |
| UNCENSORED DEATH DAYS | | | | |
| none | | | | |
| CENSORED DEATH DAYS | | | | |
| none | | | | |
| DOSE = 250 PPM | | | | |
| TOTAL 15 | UNCENSORED DEATHS 1 | CENSORED DEATHS 0 | TERMINAL 14 | |
| UNCENSORED DEATH DAYS | | | | |
| 15 | | | | |
| CENSORED DEATH DAYS | | | | |
| none | | | | |
| DOSE = 500 PPM | | | | |
| TOTAL 15 | UNCENSORED DEATHS 0 | CENSORED DEATHS 0 | TERMINAL 15 | |
| UNCENSORED DEATH DAYS | | | | |
| none | | | | |
| CENSORED DEATH DAYS | | | | |
| none | | | | |

⁽A) FIRST TERMINAL SACRIFICE

⁽B) THE FIRST ENTRY IS THE TREND TEST (TARONE, 1975) RESULT. SUBSEQUENT ENTRIES ARE THE RESULTS OF PAIRWISE TESTS (COX, 1972). NEGATIVE TRENDS ARE INDICATED BY "N".

⁽C) MEAN OF ALL UNCENSORED DEATHS PRIOR TO TERMINAL SACRIFICE

⁽D) MEAN OF ALL DEATHS (UNCENSORED, CENSORED, TERMINAL SACRIFICE)

Species/Strain: Mouse/P53(C57BL/6)

Test Type: 26-WEEK

Route: DOSED FEED

P11: STATISTICAL ANALYSIS OF SURVIVAL DATA

Test Compound: Transgenic model evaluation (Methylphenidate hydrochloride)

CAS Number: 298-59-9

Date Report Requested: 10/17/2014
Time Report Requested: 22:20:28

First Dose M/F: NA / NA

Lab: BAT

Male MOUSE FIRST TERMINAL SACRIFICE AT 169 DAYS

KAPLAN-MEIER SURVIVAL PROBABILITY ESTIMATES (%)

| DOSE | | TIME (DAYS) | | | | | | | | |
|---------|-------|-------------|-------|-------|-------|-------|-------|-------|-------|--------|
| | 17 | 34 | 51 | 68 | 85 | 102 | 119 | 136 | 153 | 169(A) |
| 0 PPM | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 50 PPM | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 250 PPM | 93.3 | 93.3 | 93.3 | 93.3 | 93.3 | 93.3 | 93.3 | 93.3 | 93.3 | 93.3 |
| 500 PPM | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |

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⁽C) MEAN OF ALL UNCENSORED DEATHS PRIOR TO TERMINAL SACRIFICE

⁽D) MEAN OF ALL DEATHS (UNCENSORED, CENSORED, TERMINAL SACRIFICE)

Species/Strain: Mouse/P53(C57BL/6)

Test Type: 26-WEEK

Route: DOSED FEED

P11: STATISTICAL ANALYSIS OF SURVIVAL DATA

Test Compound: Transgenic model evaluation (Methylphenidate hydrochloride)

CAS Number: 298-59-9

Date Report Requested: 10/17/2014
Time Report Requested: 22:20:28

First Dose M/F: NA / NA

Lab: BAT

Male MOUSE FIRST TERMINAL SACRIFICE AT 169 DAYS

SURVIVAL SUMMARY STATISTICS

| DOSE | 0 PPM | 50 PPM | 250 PPM | 500 PPM | |
|--------------------------------|---------|--------|---------|---------|--|
| SURVIVAL AT END OF STUDY | 100.0% | 100.0% | 93.3% | 100.0% | |
| (KAPLAN-MEIER) | | | | | |
| SIGNIFICANCE (B) | P=1.000 | | P=1.000 | | |
| (LIFE TABLE) | | | | | |
| MEAN DAY OF NATURAL DEATHS (C) | | | 15.0 | | |
| (STANDARD ERROR) | (.) | (.) | (.) | (.) | |
| MEAN LIFE SPAN (D) | 169.0 | 169.0 | 158.7 | 169.0 | |
| (STANDARD ERROR) | (0.0) | (0.0) | (10.3) | (0.0) | |

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Species/Strain: Mouse/P53(C57BL/6)

Test Type: 26-WEEK

Route: DOSED FEED

P11: STATISTICAL ANALYSIS OF SURVIVAL DATA

Test Compound: Transgenic model evaluation (Methylphenidate hydrochloride)

CAS Number: 298-59-9

Date Report Requested: 10/17/2014
Time Report Requested: 22:20:28

First Dose M/F: NA / NA

Lab: BAT

Female MOUSE
FIRST TERMINAL SACRIFICE AT 169 DAYS
INDIVIDUAL SURVIVAL TIMES (DAYS)

| | | <u> </u> | |
|---------------------|---|---|--|
| | | | |
| UNCENSORED DEATHS 0 | CENSORED DEATHS 0 | TERMINAL 15 | |
| | | | |
| | | | |
| | | | |
| | | | |
| | | | |
| UNCENSORED DEATHS 0 | CENSORED DEATHS 0 | TERMINAL 15 | |
| | | | |
| | | | |
| | | | |
| | | | |
| | | | |
| UNCENSORED DEATHS 0 | CENSORED DEATHS 0 | TERMINAL 15 | |
| | | | |
| | | | |
| | | | |
| | | | |
| | | | |
| UNCENSORED DEATHS 0 | CENSORED DEATHS 0 | TERMINAL 15 | |
| | | | |
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| | | | |
| | | | |
| | UNCENSORED DEATHS 0 UNCENSORED DEATHS 0 UNCENSORED DEATHS 0 | UNCENSORED DEATHS 0 CENSORED DEATHS 0 UNCENSORED DEATHS 0 CENSORED DEATHS 0 UNCENSORED DEATHS 0 CENSORED DEATHS 0 | UNCENSORED DEATHS 0 CENSORED DEATHS 0 TERMINAL 15 UNCENSORED DEATHS 0 CENSORED DEATHS 0 TERMINAL 15 |

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Species/Strain: Mouse/P53(C57BL/6)

Test Type: 26-WEEK

Route: DOSED FEED

P11: STATISTICAL ANALYSIS OF SURVIVAL DATA

Test Compound: Transgenic model evaluation (Methylphenidate hydrochloride)

CAS Number: 298-59-9

Date Report Requested: 10/17/2014 Time Report Requested: 22:20:28

First Dose M/F: NA / NA

Lab: BAT

Female MOUSE FIRST TERMINAL SACRIFICE AT 169 DAYS

KAPLAN-MEIER SURVIVAL PROBABILITY ESTIMATES (%)

| DOSE | | TIME (DAYS) | | | | | | | | |
|---------|-------|-------------|-------|-------|-------|-------|-------|-------|-------|--------|
| | 17 | 34 | 51 | 68 | 85 | 102 | 119 | 136 | 153 | 169(A) |
| 0 PPM | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 50 PPM | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 250 PPM | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |
| 500 PPM | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 | 100.0 |

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First Dose M/F: NA / NA

Lab: BAT

Female MOUSE FIRST TERMINAL SACRIFICE AT 169 DAYS

SURVIVAL SUMMARY STATISTICS

| DOSE | 0 PPM | 50 PPM | 250 PPM | 500 PPM | |
|--------------------------------|--------|--------|---------|---------|--|
| SURVIVAL AT END OF STUDY | 100.0% | 100.0% | 100.0% | 100.0% | |
| (KAPLAN-MEIER) | | | | | |
| SIGNIFICANCE (B) | | | | | |
| (LIFE TABLE) | | | | | |
| MEAN DAY OF NATURAL DEATHS (C) | • | | | | |
| (STANDARD ERROR) | (.) | (.) | (.) | (.) | |
| MEAN LIFE SPAN (D) | 169.0 | 169.0 | 169.0 | 169.0 | |
| (STANDARD ERROR) | (0.0) | (0.0) | (0.0) | (0.0) | |

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Test Type: 26-WEEK

Route: DOSED FEED

Species/Strain: Mouse/P53(C57BL/6)

P11: STATISTICAL ANALYSIS OF SURVIVAL DATA

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CAS Number: 298-59-9

Date Report Requested: 10/17/2014 Time Report Requested: 22:20:28

First Dose M/F: NA / NA

Lab: BAT

** END OF REPORT **

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⁽D) MEAN OF ALL DEATHS (UNCENSORED, CENSORED, TERMINAL SACRIFICE)