

1,3,5,7-テトラアザトリシクロ〔3.3.1.1^{3,7}〕デカンの
ラット及びマウスを用いた経口投与による
がん原性試験（混水試験）報告書

APPENDIXES

(E1～K4)

APPENDIXES (CONTINUED)

APPENDIX E 1 CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY : SUMMARY) RAT : MALE

APPENDIX E 2 CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY : SUMMARY) RAT : FEMALE

APPENDIX E 3 CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY : SUMMARY) MOUSE : MALE

APPENDIX E 4 CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY : SUMMARY) MOUSE : FEMALE

APPENDIX F 1 HEMATOLOGY(TOW-YEAR STUDY : SUMMARY) RAT MALE

APPENDIX F 2 HEMATOLOGY (TOW-YEAR STUDY : SUMMARY) RAT : FEMALE

APPENDIX F 3 HEMATOLOGY(TOW-YEAR STUDY : SUMMARY) MOUSE : MALE

APPENDIX F 4 HEMATOLOGY(TOW-YEAR STUDY : SUMMARY) MOUSE : FEMALE

APPENDIX G 1 BIOCHEMISTRY(TOW-YEAR STUDY : SUMMARY) RAT : MALE

APPENDIX G 2 BIOCHEMISTRY(TOW-YEAR STUDY : SUMMARY) RAT :FEMALE

APPENDIX G 3 BIOCHEMISTRY(TOW-YEAR STUDY : SUMMARY) MOUSE :MALE

APPENDIX G4 BIOCHEMISTRY(TOW-YEAR STUDY : SUMMARY) MOUSE :FEMALE

APPENDIX H 1 URINALYSIS (TOW-YEAR STUDY : SUMMARY) RAT : MALE

APPENDIX H 2 URINALYSIS(TOW-YEAR STUDY : SUMMARY) RAT : FEMALE

APPENDIX H 3 URINALYSIS (TOW-YEAR STUDY : SUMMARY) MOUSE : MALE

APPENDIX H 4 URINALYSIS(TOW-YEAR STUDY : SUMMARY) MOUSE : FEMALE

APPENDIXES (CONTINUED)

| | | | |
|----------|-----|---|----------------|
| APPENDIX | I 1 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) RAT : MALE : DEAD AND MORIBUND ANIMALS | |
| APPENDIX | I 2 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) RAT : FEMALE : DEAD AND MORIBUND ANIMALS | |
| APPENDIX | I 3 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) RAT : MALE : SACRIFICED ANIMALS | |
| APPENDIX | I 4 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) RAT : FEMALE : SACRIFICED ANIMALS | |
| APPENDIX | I 5 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) MOUSE : MALE : DEAD AND MORIBUND ANIMALS | |
| APPENDIX | I 6 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) MOUSE : FEMALE : DEAD AND MORIBUND ANIMALS | |
| APPENDIX | I 7 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) MOUSE : MALE : SACRIFICED ANIMALS | |
| APPENDIX | I 8 | GROSS FINDINGS (TOW-YEAR STUDY : SUMMARY) MOUSE : FEMALE : SACRIFICED ANIMALS | |
| APPENDIX | J 1 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , ABSOLUTE | RAT : MALE |
| APPENDIX | J 2 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , ABSOLUTE | RAT : FEMALE |
| APPENDIX | J 3 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , ABSOLUTE | MOUSE : MALE |
| APPENDIX | J 4 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , ABSOLUTE | MOUSE : FEMALE |
| APPENDIX | K1 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , RELATIVE | RAT : MALE |
| APPENDIX | K2 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , RELATIVE | RAT : FEMALE |
| APPENDIX | K3 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , RELATIVE | MOUSE : MALE |
| APPENDIX | K4 | ORGAN WEIGHT(TOW-YEAR STUDY : SUMMARY) , RELATIVE | MOUSE : FEMALE |

APPENDIX E 1

CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY: SUMMARY)

RAT : MALE

STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 1

| Group Name | Administration (weeks) | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.934± 0.100 | 0.849± 0.107 | 0.763± 0.037 | 0.737± 0.058 | 0.720± 0.134 | 0.649± 0.076 | 0.600± 0.043 |
| 15000 ppm | 2.002± 0.269 | 1.809± 0.429 | 1.626± 0.111 | 1.532± 0.136 | 1.436± 0.125 | 1.359± 0.109 | 1.275± 0.152 |
| 30000 ppm | 3.835± 0.365 | 3.253± 0.315 | 3.006± 0.247 | 2.883± 0.242 | 2.697± 0.206 | 2.539± 0.195 | 2.348± 0.173 |

(HAN300)

BAIS 3

STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 2

| Group Name | Administration (weeks) | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 8 | 9 | 10 | 11 | 12 | 13 | 14 |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.593± 0.054 | 0.572± 0.041 | 0.541± 0.061 | 0.516± 0.046 | 0.484± 0.035 | 0.492± 0.037 | 0.479± 0.025 |
| 15000 ppm | 1.233± 0.114 | 1.191± 0.167 | 1.116± 0.109 | 1.061± 0.081 | 0.998± 0.089 | 1.009± 0.075 | 1.016± 0.079 |
| 30000 ppm | 2.298± 0.208 | 2.244± 0.181 | 2.198± 0.231 | 2.133± 0.256 | 2.069± 0.268 | 2.041± 0.165 | 1.995± 0.130 |

(HAN300)

BAIS 3

STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 3

| Group Name | Administration (weeks) | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 16 | 18 | 20 | 22 | 24 | 26 | 28 |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.413± 0.023 | 0.401± 0.025 | 0.407± 0.044 | 0.389± 0.037 | 0.375± 0.024 | 0.369± 0.018 | 0.369± 0.032 |
| 15000 ppm | 0.859± 0.048 | 0.839± 0.100 | 0.859± 0.128 | 0.791± 0.062 | 0.803± 0.179 | 0.793± 0.072 | 0.757± 0.041 |
| 30000 ppm | 1.769± 0.130 | 1.732± 0.121 | 1.692± 0.105 | 1.622± 0.094 | 1.620± 0.132 | 1.625± 0.092 | 1.593± 0.082 |

(HAN300)

BAIS3

STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 4

| Group Name | Administration (weeks) | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 30 | 32 | 34 | 36 | 38 | 40 | 42 | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.356± 0.018 | 0.344± 0.017 | 0.322± 0.018 | 0.331± 0.016 | 0.328± 0.013 | 0.329± 0.021 | 0.328± 0.016 | |
| 15000 ppm | 0.741± 0.052 | 0.702± 0.079 | 0.672± 0.098 | 0.706± 0.103 | 0.681± 0.080 | 0.684± 0.065 | 0.687± 0.060 | |
| 30000 ppm | 1.536± 0.086 | 1.495± 0.082 | 1.383± 0.082 | 1.454± 0.111 | 1.421± 0.082 | 1.437± 0.076 | 1.434± 0.076 | |

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STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 5

| Group Name | Administration (weeks) | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 44 | 46 | 48 | 50 | 52 | 54 | 56 |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.331± 0.016 | 0.332± 0.015 | 0.334± 0.014 | 0.335± 0.018 | 0.331± 0.039 | 0.326± 0.016 | 0.324± 0.017 |
| 15000 ppm | 0.710± 0.089 | 0.703± 0.117 | 0.731± 0.122 | 0.705± 0.072 | 0.690± 0.082 | 0.704± 0.079 | 0.687± 0.099 |
| 30000 ppm | 1.491± 0.108 | 1.456± 0.083 | 1.514± 0.085 | 1.489± 0.102 | 1.471± 0.100 | 1.517± 0.084 | 1.468± 0.102 |

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STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 6

| Group Name | Administration (weeks) | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 58 | 60 | 62 | 64 | 66 | 68 | 70 | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.325± 0.019 | 0.316± 0.018 | 0.311± 0.021 | 0.315± 0.026 | 0.317± 0.020 | 0.304± 0.026 | 0.312± 0.022 | |
| 15000 ppm | 0.695± 0.103 | 0.665± 0.081 | 0.675± 0.124 | 0.681± 0.175 | 0.680± 0.094 | 0.637± 0.068 | 0.666± 0.066 | |
| 30000 ppm | 1.465± 0.098 | 1.431± 0.124 | 1.454± 0.148 | 1.439± 0.110 | 1.434± 0.113 | 1.380± 0.111 | 1.442± 0.117 | |

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BAIS 3

STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 7

| Group Name | Administration (weeks) | | 74 | 76 | 78 | 80 | 82 | 84 |
|------------|------------------------|--|--------------|--------------|--------------|--------------|--------------|--------------|
| | 72 | | | | | | | |
| Control | 0.000± 0.000 | | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.300± 0.027 | | 0.323± 0.043 | 0.320± 0.049 | 0.351± 0.056 | 0.324± 0.055 | 0.339± 0.048 | 0.340± 0.048 |
| 15000 ppm | 0.651± 0.078 | | 0.687± 0.117 | 0.694± 0.091 | 0.722± 0.084 | 0.685± 0.096 | 0.727± 0.107 | 0.736± 0.121 |
| 30000 ppm | 1.404± 0.141 | | 1.469± 0.144 | 1.447± 0.172 | 1.585± 0.197 | 1.486± 0.178 | 1.561± 0.179 | 1.603± 0.310 |

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STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 8

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 86 | 88 | 90 | 92 | 94 | 96 | 98 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 7500 ppm | 0.356± 0.057 | 0.364± 0.055 | 0.358± 0.055 | 0.388± 0.093 | 0.370± 0.071 | 0.387± 0.075 | 0.419± 0.116 | | | |
| 15000 ppm | 0.771± 0.149 | 0.756± 0.094 | 0.765± 0.140 | 0.780± 0.151 | 0.798± 0.164 | 0.874± 0.201 | 0.944± 0.278 | | | |
| 30000 ppm | 1.599± 0.205 | 1.606± 0.196 | 1.603± 0.227 | 1.703± 0.287 | 1.704± 0.306 | 1.878± 0.322 | 1.789± 0.609 | | | |

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STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 9

| Group Name | Administration (weeks) | | | | | |
|------------|------------------------|-------|--------|-------|--------|-------|
| | 100 | | 102 | | 104 | |
| Control | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 |
| 7500 ppm | 0.429± | 0.091 | 0.447± | 0.131 | 0.450± | 0.092 |
| 15000 ppm | 0.908± | 0.264 | 0.994± | 0.334 | 0.924± | 0.192 |
| 30000 ppm | 1.831± | 0.476 | 1.952± | 0.463 | 1.947± | 0.509 |

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BAIS3

APPENDIX E 2

CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE

STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 10

| Group Name | Administration (weeks) | | 2 | 3 | 4 | 5 | 6 | 7 |
|------------|------------------------|--|--------------|--------------|--------------|--------------|--------------|--------------|
| | 1 | | | | | | | |
| Control | 0.000± 0.000 | | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 1.189± 0.404 | | 1.214± 0.565 | 1.066± 0.287 | 0.965± 0.226 | 0.898± 0.209 | 0.931± 0.304 | 0.976± 0.409 |
| 15000 ppm | 2.175± 0.146 | | 2.119± 0.602 | 2.060± 0.662 | 1.919± 0.344 | 1.860± 0.555 | 1.926± 0.725 | 1.717± 0.538 |
| 30000 ppm | 4.531± 0.460 | | 4.027± 1.246 | 3.656± 0.375 | 3.629± 0.894 | 3.276± 0.320 | 3.369± 0.808 | 3.170± 0.892 |

(HAN300)

BAIS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 11

| Group Name | Administration | | (weeks) | | | | | | | | | | | |
|------------|----------------|-------|---------|-------|--------|-------|--------|-------|--------|-------|--------|-------|--------|-------|
| | 8 | | 9 | | 10 | | 11 | | 12 | | 13 | | 14 | |
| Control | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 |
| 7500 ppm | 0.905± | 0.279 | 0.903± | 0.285 | 0.915± | 0.333 | 0.846± | 0.309 | 0.792± | 0.253 | 0.828± | 0.282 | 0.825± | 0.210 |
| 15000 ppm | 1.749± | 0.572 | 1.708± | 0.473 | 1.584± | 0.308 | 1.645± | 0.466 | 1.559± | 0.548 | 1.438± | 0.267 | 1.679± | 0.520 |
| 30000 ppm | 3.282± | 1.186 | 3.042± | 0.701 | 3.002± | 0.805 | 2.762± | 0.614 | 3.104± | 1.295 | 2.937± | 0.811 | 3.031± | 0.883 |

(HAN300)

BAIS3

STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 12

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 16 | 18 | 20 | 22 | 24 | 26 | 28 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 7500 ppm | 0.761± 0.249 | 0.722± 0.176 | 0.705± 0.175 | 0.725± 0.258 | 0.698± 0.181 | 0.723± 0.255 | 0.645± 0.168 | | | |
| 15000 ppm | 1.545± 0.557 | 1.533± 0.417 | 1.477± 0.402 | 1.381± 0.366 | 1.445± 0.436 | 1.414± 0.414 | 1.319± 0.373 | | | |
| 30000 ppm | 2.594± 0.547 | 2.634± 0.428 | 2.656± 0.531 | 2.580± 0.739 | 2.690± 0.791 | 2.699± 0.738 | 2.530± 0.615 | | | |

(HAN300)

BAIS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 13

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 30 | 32 | 34 | 36 | 38 | 40 | 42 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 7500 ppm | 0.655± 0.189 | 0.616± 0.191 | 0.546± 0.153 | 0.674± 0.296 | 0.582± 0.172 | 0.594± 0.186 | 0.617± 0.239 | | | |
| 15000 ppm | 1.249± 0.310 | 1.375± 0.550 | 1.123± 0.320 | 1.247± 0.384 | 1.267± 0.451 | 1.195± 0.398 | 1.194± 0.320 | | | |
| 30000 ppm | 2.379± 0.523 | 2.613± 0.789 | 2.161± 0.309 | 2.399± 0.612 | 2.317± 0.635 | 2.302± 0.567 | 2.343± 0.686 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 14

| Group Name | Administration (weeks) | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 44 | 46 | 48 | 50 | 52 | 54 | 56 | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.573± 0.164 | 0.551± 0.136 | 0.562± 0.113 | 0.593± 0.199 | 0.586± 0.194 | 0.542± 0.142 | 0.531± 0.161 | |
| 15000 ppm | 1.154± 0.240 | 1.167± 0.270 | 1.221± 0.363 | 1.231± 0.407 | 1.163± 0.323 | 1.056± 0.162 | 1.124± 0.321 | |
| 30000 ppm | 2.507± 0.803 | 2.457± 0.508 | 2.419± 0.542 | 2.447± 0.535 | 2.504± 0.755 | 2.385± 0.353 | 2.437± 0.637 | |

(HAN300)

BATS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 15

| Group Name | Administration (weeks) | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|
| | 58 | 60 | 62 | 64 | 66 | 68 | 70 | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.552± 0.134 | 0.508± 0.108 | 0.536± 0.160 | 0.511± 0.149 | 0.507± 0.179 | 0.457± 0.128 | 0.440± 0.123 | |
| 15000 ppm | 1.154± 0.299 | 1.118± 0.314 | 1.046± 0.209 | 1.021± 0.284 | 0.984± 0.209 | 0.947± 0.226 | 0.834± 0.208 | |
| 30000 ppm | 2.309± 0.414 | 2.360± 0.649 | 2.251± 0.709 | 2.202± 0.431 | 2.297± 0.669 | 2.112± 0.382 | 2.143± 0.456 | |

(HAN300)

BAIS 3

STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 16

| Group Name | Administration (weeks) | | 74 | 76 | 78 | 80 | 82 | 84 |
|------------|------------------------|--|--------------|--------------|--------------|--------------|--------------|--------------|
| | 72 | | | | | | | |
| Control | 0.000± 0.000 | | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.447± 0.110 | | 0.465± 0.106 | 0.448± 0.096 | 0.458± 0.092 | 0.449± 0.124 | 0.483± 0.141 | 0.470± 0.135 |
| 15000 ppm | 0.972± 0.373 | | 0.965± 0.289 | 0.982± 0.250 | 1.031± 0.250 | 0.960± 0.315 | 1.035± 0.311 | 0.954± 0.205 |
| 30000 ppm | 2.191± 0.619 | | 2.203± 0.597 | 2.146± 0.454 | 2.167± 0.635 | 2.165± 0.686 | 2.217± 0.363 | 2.343± 0.746 |

(HAN300)

BAIS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 17

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 86 | 88 | 90 | 92 | 94 | 96 | 98 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 7500 ppm | 0.478± 0.093 | 0.470± 0.108 | 0.456± 0.103 | 0.484± 0.126 | 0.450± 0.104 | 0.502± 0.114 | 0.484± 0.130 | | | |
| 15000 ppm | 0.989± 0.188 | 0.982± 0.213 | 0.958± 0.179 | 1.025± 0.211 | 0.976± 0.227 | 1.024± 0.259 | 1.023± 0.230 | | | |
| 30000 ppm | 2.251± 0.526 | 2.289± 0.462 | 2.224± 0.430 | 2.344± 0.649 | 2.282± 0.537 | 2.466± 0.722 | 2.477± 0.853 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0224
ANIMAL : RAT F344
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 18

| Group Name | Administration (weeks) | | |
|------------|------------------------|--------------|--------------|
| | 100 | 102 | 104 |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 7500 ppm | 0.507± 0.131 | 0.558± 0.185 | 0.557± 0.154 |
| 15000 ppm | 1.070± 0.301 | 1.127± 0.251 | 1.102± 0.231 |
| 30000 ppm | 2.257± 0.656 | 2.130± 0.516 | 2.380± 0.515 |

(HAN300)

BAIS3

APPENDIX E 3

CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 1

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|---------------|---------------|---------------|---------------|--------------|--|--|
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 2.235± 0.281 | 2.047± 0.326 | 1.936± 0.245 | 2.056± 0.346 | 1.900± 0.555 | 2.222± 0.864 | 2.222± 0.871 | | | |
| 20000 ppm | 4.360± 0.560 | 4.185± 0.502 | 4.047± 0.658 | 4.420± 0.664 | 4.078± 0.905 | 4.699± 1.322 | 4.829± 2.014 | | | |
| 40000 ppm | 9.635± 1.380 | 9.219± 1.191 | 8.968± 1.164 | 10.666± 1.720 | 10.688± 5.269 | 10.374± 3.426 | 11.089± 5.203 | | | |

(HAN300)

BAIS3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/d a y
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 2

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 8 | 9 | 10 | 11 | 12 | 13 | 14 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 2.138± 1.062 | 2.021± 0.395 | 1.812± 0.563 | 1.690± 0.401 | 1.755± 0.486 | 1.622± 0.601 | 1.552± 0.316 | | | |
| 20000 ppm | 4.696± 1.772 | 4.438± 1.144 | 3.831± 1.187 | 4.092± 2.270 | 3.543± 0.639 | 3.327± 1.495 | 3.502± 1.623 | | | |
| 40000 ppm | 10.460± 4.137 | 9.507± 2.928 | 8.374± 2.160 | 8.501± 3.427 | 8.005± 1.733 | 7.406± 2.348 | 7.420± 2.788 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 3

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 16 | 18 | 20 | 22 | 24 | 26 | 28 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 1.561± 0.633 | 1.291± 0.179 | 1.161± 0.235 | 1.171± 0.199 | 1.164± 0.480 | 1.120± 0.201 | 1.038± 0.179 | | | |
| 20000 ppm | 3.057± 1.140 | 2.830± 0.945 | 2.513± 0.617 | 2.492± 0.644 | 2.136± 0.328 | 2.126± 0.229 | 2.232± 0.835 | | | |
| 40000 ppm | 6.802± 2.157 | 6.215± 1.814 | 5.623± 1.574 | 5.792± 1.282 | 5.342± 1.824 | 4.776± 0.590 | 4.761± 0.544 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 4

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 30 | 32 | 34 | 36 | 38 | 40 | 42 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 1.011± 0.210 | 0.959± 0.139 | 0.977± 0.165 | 1.009± 0.168 | 0.905± 0.103 | 0.941± 0.119 | 0.973± 0.168 | | | |
| 20000 ppm | 2.058± 0.681 | 2.014± 0.534 | 2.094± 1.396 | 1.945± 0.593 | 1.959± 0.707 | 2.052± 0.559 | 2.025± 0.381 | | | |
| 40000 ppm | 4.427± 1.139 | 4.663± 0.944 | 4.436± 0.705 | 4.313± 0.568 | 4.117± 0.538 | 4.469± 0.665 | 4.289± 0.782 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 5

| Group Name | Administration (weeks) | | | | | | | | | | | | | |
|------------|------------------------|-------|--------|-------|--------|-------|--------|-------|--------|-------|--------|-------|--------|-------|
| | 44 | | 46 | | 48 | | 50 | | 52 | | 54 | | 56 | |
| Control | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 | 0.000± | 0.000 |
| 10000 ppm | 0.948± | 0.098 | 0.927± | 0.103 | 0.974± | 0.108 | 0.951± | 0.101 | 0.865± | 0.146 | 0.923± | 0.091 | 0.942± | 0.114 |
| 20000 ppm | 2.024± | 0.582 | 1.930± | 0.472 | 2.001± | 0.235 | 1.895± | 0.351 | 1.767± | 0.313 | 1.911± | 0.284 | 2.003± | 0.386 |
| 40000 ppm | 4.509± | 1.323 | 4.365± | 0.572 | 4.637± | 1.104 | 4.324± | 1.467 | 4.113± | 1.238 | 4.636± | 2.824 | 4.867± | 2.800 |

(HAN300)

BAIS 3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 6

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 58 | 60 | 62 | 64 | 66 | 68 | 70 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 0.924± 0.104 | 0.915± 0.148 | 0.890± 0.113 | 1.092± 1.172 | 0.990± 0.357 | 0.877± 0.096 | 0.847± 0.068 | | | |
| 20000 ppm | 2.008± 0.254 | 1.938± 0.247 | 1.756± 0.199 | 1.860± 0.258 | 1.899± 0.272 | 1.892± 0.315 | 1.765± 0.256 | | | |
| 40000 ppm | 4.690± 2.203 | 4.524± 2.520 | 3.948± 0.544 | 4.232± 0.796 | 4.321± 1.585 | 4.531± 2.443 | 4.320± 2.256 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 7

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 72 | 74 | 76 | 78 | 80 | 82 | 84 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 0.846± 0.059 | 0.869± 0.120 | 0.853± 0.103 | 0.849± 0.081 | 0.879± 0.106 | 0.868± 0.155 | 0.902± 0.270 | | | |
| 20000 ppm | 1.860± 0.376 | 1.835± 0.349 | 1.934± 0.541 | 1.871± 0.340 | 1.927± 0.465 | 1.915± 0.594 | 1.975± 0.783 | | | |
| 40000 ppm | 4.687± 3.404 | 5.002± 3.994 | 5.186± 4.606 | 4.910± 4.782 | 4.503± 2.194 | 4.679± 2.824 | 4.879± 3.355 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 8

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 86 | 88 | 90 | 92 | 94 | 96 | 98 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 0.964± 0.578 | 0.947± 0.318 | 0.966± 0.576 | 0.930± 0.135 | 0.895± 0.159 | 0.983± 0.321 | 0.926± 0.216 | | | |
| 20000 ppm | 2.078± 0.824 | 2.298± 1.061 | 2.766± 1.906 | 2.722± 1.830 | 2.913± 2.614 | 3.211± 2.992 | 3.027± 2.831 | | | |
| 40000 ppm | 5.394± 4.141 | 5.877± 5.244 | 6.762± 8.220 | 5.971± 5.382 | 6.749± 7.050 | 6.981± 7.286 | 8.180± 8.426 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : MALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

| Group Name | Administration (weeks) | | |
|------------|------------------------|--------------|--------------|
| | 100 | 102 | 104 |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 10000 ppm | 0.955± 0.260 | 1.025± 0.358 | 1.015± 0.187 |
| 20000 ppm | 2.712± 1.199 | 2.577± 1.084 | 2.779± 1.036 |
| 40000 ppm | 8.131± 8.329 | 8.380± 8.959 | 8.363± 9.097 |

APPENDIX E 4

CHEMICAL INTAKE CHANGES (TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/d a y
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 10

| Group Name | Administration (weeks) | | 2 | 3 | 4 | 5 | 6 | 7 |
|------------|------------------------|--|---------------|---------------|---------------|---------------|---------------|---------------|
| | 1 | | | | | | | |
| Control | 0.000± 0.000 | | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 |
| 10000 ppm | 2.521± 0.204 | | 2.537± 0.305 | 2.373± 0.328 | 2.535± 0.531 | 2.918± 1.318 | 3.308± 1.720 | 3.678± 2.357 |
| 20000 ppm | 5.150± 0.749 | | 5.405± 0.811 | 5.264± 0.706 | 6.740± 2.051 | 6.988± 4.085 | 7.265± 4.373 | 6.559± 2.966 |
| 40000 ppm | 11.587± 1.192 | | 11.502± 1.295 | 10.920± 1.348 | 14.617± 6.218 | 12.515± 5.566 | 13.930± 5.465 | 14.238± 5.857 |

(HAN300)

BAIS 3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
UNIT : g/kg/d a y
REPORT TYPE : A1 104
SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 11

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|---------------|---------------|---------------|---------------|----------------|---------------|--------------|--|--|
| | 8 | 9 | 10 | 11 | 12 | 13 | 14 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 3.856± 3.248 | 3.412± 1.699 | 3.131± 2.590 | 3.394± 1.902 | 3.321± 1.631 | 3.234± 2.456 | 3.856± 2.637 | | | |
| 20000 ppm | 6.569± 2.801 | 6.262± 3.137 | 6.330± 4.155 | 7.335± 6.016 | 6.425± 3.994 | 7.861± 6.765 | 7.595± 5.170 | | | |
| 40000 ppm | 13.957± 7.365 | 12.870± 7.587 | 11.378± 5.405 | 11.916± 4.065 | 12.008± 5.177 | 13.406± 11.439 | 13.541± 8.382 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 12

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|---------------|---------------|--------------|---------------|--------------|--------------|--------------|--|--|
| | 16 | 18 | 20 | 22 | 24 | 26 | 28 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 3.133± 1.789 | 2.878± 1.413 | 2.529± 1.149 | 2.547± 1.191 | 2.403± 1.230 | 2.187± 0.894 | 2.208± 1.065 | | | |
| 20000 ppm | 5.807± 3.009 | 5.879± 3.041 | 4.646± 1.834 | 4.675± 1.654 | 4.522± 1.313 | 4.748± 1.580 | 4.403± 1.615 | | | |
| 40000 ppm | 11.183± 3.488 | 11.207± 5.640 | 10.177± 6.637 | 9.126± 3.580 | 10.404± 6.032 | 8.996± 2.949 | 9.258± 4.361 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 13

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 30 | 32 | 34 | 36 | 38 | 40 | 42 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 2.384± 1.056 | 1.923± 0.681 | 1.854± 0.885 | 2.008± 0.781 | 1.920± 1.012 | 1.917± 0.941 | 2.186± 1.307 | | | |
| 20000 ppm | 4.188± 1.528 | 4.033± 1.283 | 4.409± 2.868 | 4.256± 1.940 | 3.955± 1.525 | 4.281± 2.321 | 4.291± 2.285 | | | |
| 40000 ppm | 8.666± 2.790 | 9.386± 5.476 | 8.296± 2.694 | 8.129± 4.638 | 7.321± 3.173 | 7.767± 2.457 | 7.306± 1.701 | | | |

(HAN300)

BAIS 3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/d a y
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 14

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 44 | 46 | 48 | 50 | 52 | 54 | 56 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 2.118± 1.262 | 1.969± 1.388 | 1.992± 1.099 | 1.993± 0.986 | 1.708± 0.763 | 2.271± 1.820 | 1.974± 1.058 | | | |
| 20000 ppm | 4.545± 2.926 | 4.008± 3.342 | 3.697± 1.097 | 4.123± 3.463 | 3.507± 1.673 | 3.830± 1.887 | 3.733± 1.147 | | | |
| 40000 ppm | 7.396± 1.640 | 7.385± 2.061 | 7.558± 1.675 | 7.681± 2.975 | 6.639± 1.383 | 7.005± 1.867 | 7.118± 1.348 | | | |

(HAN300)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 15

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 58 | 60 | 62 | 64 | 66 | 68 | 70 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 1.724± 0.741 | 1.758± 0.834 | 1.494± 0.618 | 1.552± 0.742 | 1.484± 0.647 | 1.451± 0.667 | 1.320± 0.557 | | | |
| 20000 ppm | 3.478± 1.677 | 3.541± 1.983 | 2.957± 1.170 | 3.310± 1.514 | 3.468± 3.265 | 3.336± 2.253 | 2.914± 1.219 | | | |
| 40000 ppm | 6.273± 1.106 | 6.836± 2.428 | 5.664± 0.899 | 6.185± 1.470 | 5.676± 0.800 | 5.751± 0.912 | 5.669± 1.346 | | | |

(HAN300)

BAIS3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 16

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 72 | 74 | 76 | 78 | 80 | 82 | 84 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 1.344± 0.481 | 1.431± 0.918 | 1.373± 0.723 | 1.157± 0.360 | 1.319± 0.631 | 1.254± 0.536 | 1.259± 0.518 | | | |
| 20000 ppm | 3.076± 1.399 | 2.889± 0.882 | 2.870± 1.082 | 2.674± 0.997 | 2.953± 1.338 | 2.675± 0.596 | 2.688± 0.719 | | | |
| 40000 ppm | 5.785± 1.008 | 5.879± 1.005 | 5.552± 0.895 | 5.382± 0.994 | 5.867± 1.475 | 5.690± 1.627 | 5.579± 1.048 | | | |

(HAN300)

BAIS3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 UNIT : g/kg/day
 REPORT TYPE : A1 104
 SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
 ALL ANIMALS

PAGE : 17

| Group Name | Administration (weeks) | | | | | | | | | |
|------------|------------------------|--------------|--------------|--------------|--------------|--------------|--------------|--------------|--|--|
| | 86 | 88 | 90 | 92 | 94 | 96 | 98 | | | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | | |
| 10000 ppm | 1.251± 0.331 | 1.326± 0.370 | 1.250± 0.235 | 1.286± 0.342 | 1.253± 0.365 | 1.352± 0.413 | 1.271± 0.283 | | | |
| 20000 ppm | 2.762± 0.741 | 2.887± 0.884 | 2.957± 0.749 | 2.964± 0.742 | 2.915± 0.833 | 3.110± 0.800 | 3.143± 1.148 | | | |
| 40000 ppm | 5.580± 1.018 | 5.909± 1.316 | 5.842± 1.287 | 6.226± 2.249 | 6.501± 3.015 | 7.236± 4.991 | 7.011± 4.289 | | | |

(HAN300)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
UNIT : g/kg/day
REPORT TYPE : A1 104
SEX : FEMALE

CHEMICAL INTAKE CHANGES (SUMMARY)
ALL ANIMALS

PAGE : 18

| Group Name | Administration (weeks) | | | |
|------------|------------------------|--------------|--------------|--|
| | 100 | 102 | 104 | |
| Control | 0.000± 0.000 | 0.000± 0.000 | 0.000± 0.000 | |
| 10000 ppm | 1.402± 0.281 | 1.314± 0.205 | 1.340± 0.284 | |
| 20000 ppm | 3.702± 2.232 | 4.014± 3.627 | 3.062± 0.485 | |
| 40000 ppm | 6.594± 2.595 | 6.454± 2.584 | 7.502± 4.618 | |

(HAN300)

BAIS3

APPENDIX F 1

HEMATOLOGY (TOW-YEAR STUDY: SUMMARY)

RAT : MALE

STUDY NO. : 0224
ANIMAL : RAT F344
SAMPLING DATE : 105-1
SEX : MALE

HEMATOLOGY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of Animals | RED BLOOD CELL 10 ⁶ /μl | | HEMOGLOBIN g/dl | | HEMATOCRIT % | | MCV fl | | MCH pg | | MCHC g/dl | | PLATELET 10 ⁹ /μl | |
|------------|-------------------|---------------------------------------|------|--------------------|-----|-----------------|-----|-----------|-----|-----------|-----|--------------|-----|---------------------------------|-------|
| Control | 38 | 8.12± | 1.51 | 14.5± | 2.7 | 43.2± | 7.5 | 53.4± | 3.0 | 17.8± | 1.3 | 33.4± | 1.7 | 881± | 178 |
| 7500 ppm | 39 | 8.48± | 1.73 | 15.0± | 2.8 | 44.7± | 7.5 | 53.8± | 8.3 | 18.0± | 2.0 | 33.5± | 1.9 | 865± | 232 |
| 15000 ppm | 34 | 8.62± | 1.19 | 15.3± | 2.0 | 45.4± | 5.3 | 52.9± | 2.3 | 17.8± | 0.5 | 33.7± | 1.1 | 880± | 109 |
| 30000 ppm | 29 | 8.83± | 1.74 | 15.9± | 2.6 | 47.0± | 7.4 | 54.2± | 6.9 | 18.3± | 2.4 | 33.7± | 1.3 | 747± | 167** |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-1
 SEX : MALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of Animals | WBC 10 ³ /μl | | Differential N-BAND | | WBC (%) N-SEG | | EOSINO | | BASO | | MONO | | LYMPHO | | OTHERS | |
|------------|-------------------|----------------------------|------|------------------------|---|------------------|----|--------|---|------|---|------|---|--------|----|--------|----|
| Control | 38 | 5.56± | 1.27 | 1± | 1 | 58± | 10 | 2± | 2 | 0± | 0 | 4± | 2 | 32± | 9 | 3± | 3 |
| 7500 ppm | 39 | 8.11± | 9.83 | 1± | 1 | 55± | 12 | 1± | 1 | 0± | 0 | 4± | 2 | 34± | 11 | 6± | 13 |
| 15000 ppm | 34 | 5.86± | 2.49 | 1± | 2 | 55± | 11 | 2± | 1 | 0± | 0 | 4± | 2 | 35± | 8 | 3± | 4 |
| 30000 ppm | 29 | 5.59± | 1.60 | 1± | 2 | 53± | 9 | 2± | 1 | 0± | 0 | 4± | 1 | 36± | 9 | 4± | 8 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

APPENDIX F 2

HEMATOLOGY (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-1
 SEX : FEMALE

REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | RED BLOOD CELL 10 ⁶ /μl | | HEMOGLOBIN g/dl | | HEMATOCRIT % | | MCV fl | | MCH pg | | MCHC g/dl | | PLATELET 10 ³ /μl | |
|------------|-------------------|---------------------------------------|------|--------------------|-----|-----------------|-----|-----------|-----|-----------|-----|--------------|-----|---------------------------------|------|
| Control | 40 | 7.91± | 0.71 | 14.8± | 1.5 | 43.1± | 4.3 | 54.5± | 2.9 | 18.7± | 1.1 | 34.4± | 1.3 | 684± | 95 |
| 7500 ppm | 34 | 7.82± | 1.02 | 14.7± | 1.9 | 43.5± | 4.1 | 56.2± | 4.8 | 18.8± | 0.9 | 33.6± | 2.1 | 632± | 104 |
| 15000 ppm | 31 | 7.80± | 1.24 | 14.8± | 2.1 | 43.3± | 6.0 | 56.3± | 6.5 | 19.1± | 1.7 | 34.1± | 1.6 | 643± | 112 |
| 30000 ppm | 11 | 7.80± | 0.95 | 14.6± | 1.3 | 43.3± | 3.2 | 56.0± | 4.4 | 18.8± | 1.0 | 33.6± | 0.8 | 590± | 123* |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-1
 SEX : FEMALE

REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | WBC 10 ³ /μl | | Differential N-BAND | | WBC (%) N-SEG | | EOSINO | | BASO | | MONO | | LYMPHO | | OTHERS | |
|------------|-------------------|----------------------------|-------|------------------------|---|------------------|----|--------|---|------|---|------|---|--------|----|--------|----|
| Control | 40 | 2.82± | 1.04 | 1± | 2 | 53± | 9 | 2± | 1 | 0± | 0 | 4± | 2 | 37± | 9 | 3± | 2 |
| 7500 ppm | 34 | 8.89± | 29.44 | 1± | 1 | 50± | 17 | 2± | 1 | 0± | 0 | 4± | 2 | 35± | 12 | 8± | 20 |
| 15000 ppm | 31 | 4.08± | 4.72 | 1± | 1 | 50± | 15 | 1± | 1 | 0± | 0 | 4± | 2 | 37± | 14 | 6± | 16 |
| 30000 ppm | 11 | 3.31± | 2.21 | 1± | 1 | 47± | 12 | 1± | 1 | 0± | 0 | 4± | 2 | 39± | 9 | 8± | 11 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

APPENDIX F 3

HEMATOLOGY (TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 SAMPLING DATE : 105-1
 SEX : MALE

REPORT TYPE : A1

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of Animals | RED BLOOD CELL 10 ⁶ /μl | | HEMOGLOBIN g/dl | | HEMATOCRIT % | | MCV fl | | MCH pg | | MCHC g/dl | | PLATELET 10 ³ /μl | |
|------------|-------------------|---------------------------------------|------|--------------------|-----|-----------------|-----|-----------|-----|-----------|-----|--------------|-----|---------------------------------|-----|
| Control | 31 | 9.57± | 1.18 | 13.4± | 1.5 | 41.2± | 4.5 | 43.3± | 2.6 | 14.1± | 0.9 | 32.5± | 1.0 | 1857± | 498 |
| 10000 ppm | 35 | 9.57± | 1.07 | 13.6± | 1.5 | 41.8± | 4.1 | 43.8± | 2.1 | 14.2± | 0.7 | 32.5± | 1.5 | 1854± | 506 |
| 20000 ppm | 27 | 9.59± | 1.85 | 13.2± | 2.3 | 40.9± | 7.1 | 42.9± | 3.6 | 13.8± | 1.0 | 32.2± | 1.5 | 1933± | 531 |
| 40000 ppm | 25 | 9.57± | 1.10 | 13.4± | 1.9 | 41.3± | 4.9 | 43.2± | 1.8 | 13.9± | 0.7 | 32.3± | 1.5 | 1957± | 457 |

Significant difference : * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 SAMPLING DATE : 105-1
 SEX : MALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of Animals | WBC 10 ³ /μl | | Differential N-BAND | | WBC (%) N-SEG | | EOSINO | | BASO | | MONO | | LYMPHO | | OTHERS | |
|------------|-------------------|----------------------------|------|------------------------|---|------------------|----|--------|---|------|---|------|---|--------|----|--------|----|
| Control | 31 | 2.97± | 1.57 | 0± | 1 | 32± | 14 | 2± | 2 | 0± | 0 | 4± | 2 | 61± | 14 | 0± | 1 |
| 10000 ppm | 35 | 3.27± | 1.90 | 0± | 1 | 28± | 13 | 1± | 1 | 0± | 0 | 3± | 2 | 67± | 13 | 1± | 1 |
| 20000 ppm | 27 | 3.96± | 5.80 | 1± | 1 | 32± | 17 | 1± | 1 | 0± | 0 | 4± | 2 | 60± | 18 | 3± | 11 |
| 40000 ppm | 25 | 2.26± | 0.99 | 0± | 1 | 37± | 17 | 1± | 1 | 0± | 0 | 4± | 2 | 57± | 18 | 1± | 1 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

APPENDIX F 4

HEMATOLOGY (TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 SAMPLING DATE : 105-1
 SEX : FEMALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 3

| Group Name | NO. of Animals | RED BLOOD CELL 10 ⁶ /μl | | HEMOGLOBIN g/dl | | HEMATOCRIT % | | MCV fl | | MCH pg | | MCHC g/dl | | PLATELET 10 ³ /μl | |
|------------|-------------------|---------------------------------------|------|--------------------|-----|-----------------|-----|-----------|-----|-----------|-----|--------------|-----|---------------------------------|-----|
| Control | 28 | 9.58± | 0.66 | 14.0± | 1.0 | 42.7± | 2.7 | 44.6± | 1.5 | 14.6± | 0.5 | 32.8± | 0.7 | 1173± | 209 |
| 10000 ppm | 23 | 9.76± | 0.52 | 14.2± | 0.8 | 43.3± | 2.4 | 44.4± | 1.0 | 14.6± | 0.6 | 32.8± | 1.2 | 1098± | 182 |
| 20000 ppm | 17 | 8.93± | 1.46 | 13.2± | 2.2 | 40.4± | 5.5 | 45.6± | 3.3 | 14.7± | 0.6 | 32.4± | 1.7 | 1083± | 253 |
| 40000 ppm | 26 | 9.53± | 0.86 | 13.9± | 1.3 | 42.5± | 3.4 | 44.7± | 1.6 | 14.6± | 0.6 | 32.7± | 0.8 | 1089± | 309 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 SAMPLING DATE : 105-1
 SEX : FEMALE

HEMATOLOGY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 4

| Group Name | NO. of Animals | WBC 10 ³ /μl | | Differential N-BAND | | WBC (%) N-SEG | | EOSINO | | BASO | | MONO | | LYMPHO | | OTHERS | |
|------------|-------------------|----------------------------|------|------------------------|---|------------------|----|--------|---|------|---|------|---|--------|----|--------|---|
| Control | 28 | 1.83± | 1.38 | 0± | 1 | 29± | 12 | 2± | 2 | 0± | 0 | 3± | 2 | 64± | 13 | 1± | 2 |
| 10000 ppm | 23 | 1.71± | 1.01 | 1± | 1 | 26± | 11 | 2± | 2 | 0± | 0 | 4± | 2 | 67± | 12 | 0± | 1 |
| 20000 ppm | 17 | 2.05± | 0.99 | 1± | 1 | 28± | 14 | 1± | 2 | 0± | 0 | 4± | 2 | 64± | 15 | 2± | 3 |
| 40000 ppm | 26 | 3.25± | 3.17 | 1± | 1 | 24± | 11 | 2± | 3 | 0± | 0 | 4± | 2 | 68± | 12 | 2± | 2 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL070)

BAIS3

APPENDIX G 1

BIOCHEMISTRY (TOW-YEAR STUDY: SUMMARY)

RAT : MALE

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-2
 SEX : MALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of Animals | TOTAL PROTEIN g/dl | | ALBUMIN g/dl | | A/G RATIO | | T-BILIRUBIN mg/dl | | GLUCOSE mg/dl | | T-CHOLESTEROL mg/dl | | TRIGLYCERIDE mg/dl | |
|------------|-------------------|-----------------------|-------|-----------------|-----|-----------|-------|----------------------|------|------------------|----|------------------------|------|-----------------------|------|
| Control | 38 | 6.5± | 0.4 | 3.2± | 0.2 | 1.0± | 0.1 | 0.24± | 0.03 | 152± | 21 | 227± | 77 | 258± | 161 |
| 7500 ppm | 39 | 6.5± | 0.5 | 3.2± | 0.3 | 1.0± | 0.1 | 0.27± | 0.13 | 152± | 25 | 211± | 71 | 245± | 195 |
| 15000 ppm | 34 | 6.5± | 0.4 | 3.2± | 0.2 | 1.0± | 0.1 | 0.24± | 0.02 | 156± | 35 | 210± | 57 | 225± | 122 |
| 30000 ppm | 29 | 6.2± | 0.4** | 3.2± | 0.2 | 1.1± | 0.1** | 0.25± | 0.07 | 147± | 17 | 140± | 35** | 122± | 45** |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

STUDY NO. : 0224
ANIMAL : RAT F344
SAMPLING DATE : 105-2
SEX : MALE

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of Animals | PHOSPHOLIPID mg/dl | | GOT IU/l | | GPT IU/l | | LDH IU/l | | ALP IU/l | | G-GTP IU/l | | CPK IU/l | |
|------------|-------------------|-----------------------|------|-------------|------|-------------|----|-------------|-------|-------------|-----|---------------|---|-------------|----|
| Control | 38 | 316± | 100 | 70± | 58 | 34± | 20 | 163± | 41 | 157± | 55 | 7± | 4 | 81± | 12 |
| 7500 ppm | 39 | 297± | 97 | 84± | 87 | 38± | 25 | 230± | 281* | 179± | 110 | 8± | 6 | 88± | 35 |
| 15000 ppm | 34 | 289± | 69 | 66± | 23 | 30± | 7 | 178± | 40 | 143± | 33 | 7± | 4 | 80± | 12 |
| 30000 ppm | 29 | 202± | 42** | 87± | 41** | 38± | 19 | 226± | 152** | 158± | 60 | 7± | 6 | 107± | 92 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-2
 SEX : MALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | UREA NITROGEN mg/dl | | CREATININE mg/dl | | SODIUM mEq/l | | POTASSIUM mEq/l | | CHLORIDE mEq/l | | CALCIUM mg/dl | | INORGANIC PHOSPHORUS mg/dl | |
|------------|-------------------|------------------------|------|---------------------|-----|-----------------|---|--------------------|-----|-------------------|----|------------------|-------|-------------------------------|-------|
| Control | 38 | 27.4± | 24.0 | 0.9± | 0.6 | 144± | 3 | 3.8± | 0.3 | 107± | 2 | 11.3± | 0.8 | 5.0± | 1.9 |
| 7500 ppm | 39 | 29.0± | 30.8 | 0.8± | 0.2 | 144± | 3 | 3.9± | 0.4 | 107± | 2 | 11.4± | 0.9 | 5.2± | 2.6 |
| 15000 ppm | 34 | 25.2± | 7.0 | 0.8± | 0.2 | 144± | 2 | 3.9± | 0.4 | 108± | 2 | 11.1± | 0.4 | 4.4± | 0.8 |
| 30000 ppm | 29 | 38.0± | 74.2 | 1.0± | 1.6 | 145± | 5 | 4.0± | 0.8 | 109± | 2* | 10.8± | 0.3** | 4.8± | 4.9** |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

APPENDIX G 2

BIOCHEMISTRY (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-2
 SEX : FEMALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | TOTAL PROTEIN g/dl | | ALBUMIN g/dl | | A/G RATIO | | T-BILIRUBIN mg/dl | | GLUCOSE mg/dl | | T-CHOLESTEROL mg/dl | | TRIGLYCERIDE mg/dl | |
|------------|-------------------|-----------------------|-----|-----------------|-----|-----------|-----|----------------------|------|------------------|----|------------------------|----|-----------------------|-----|
| Control | 40 | 6.8± | 0.5 | 3.7± | 0.3 | 1.2± | 0.1 | 0.21± | 0.05 | 152± | 19 | 163± | 64 | 141± | 154 |
| 7500 ppm | 34 | 6.9± | 0.3 | 3.7± | 0.2 | 1.2± | 0.1 | 0.24± | 0.07 | 151± | 21 | 162± | 53 | 166± | 155 |
| 15000 ppm | 31 | 6.8± | 0.4 | 3.6± | 0.2 | 1.2± | 0.1 | 0.31± | 0.58 | 147± | 20 | 167± | 49 | 149± | 131 |
| 30000 ppm | 11 | 6.6± | 0.3 | 3.6± | 0.2 | 1.2± | 0.1 | 0.26± | 0.20 | 156± | 26 | 144± | 19 | 77± | 32 |

Significant defference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-2
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 5

| Group Name | NO. of Animals | PHOSPHOLIPID mg/dl | | GOT IU/l | | GPT IU/l | | LDH IU/l | | ALP IU/l | | G-GTP IU/l | | CPK IU/l | |
|------------|-------------------|-----------------------|----|-------------|-------|-------------|------|-------------|-----|-------------|----|---------------|---|-------------|----|
| Control | 40 | 274± | 93 | 97± | 48 | 46± | 17 | 189± | 67 | 116± | 42 | 3± | 2 | 80± | 16 |
| 7500 ppm | 34 | 272± | 80 | 117± | 84 | 53± | 24 | 256± | 219 | 123± | 52 | 3± | 2 | 86± | 41 |
| 15000 ppm | 31 | 284± | 85 | 164± | 302 | 65± | 59 | 328± | 574 | 130± | 88 | 4± | 5 | 93± | 68 |
| 30000 ppm | 11 | 240± | 32 | 192± | 175** | 79± | 48** | 215± | 47 | 154± | 84 | 6± | 5 | 93± | 31 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS 3

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 105-2
 SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 6

| Group Name | NO. of Animals | UREA NITROGEN mg/dl | | CREATININE mg/dl | | SODIUM mEq/l | | POTASSIUM mEq/l | | CHLORIDE mEq/l | | CALCIUM mg/dl | | INORGANIC PHOSPHORUS mg/dl | |
|------------|-------------------|------------------------|------|---------------------|-----|-----------------|---|--------------------|------|-------------------|---|------------------|-----|-------------------------------|-----|
| Control | 40 | 16.2± | 2.0 | 0.6± | 0.1 | 142± | 2 | 3.8± | 0.4 | 107± | 2 | 10.9± | 0.4 | 3.8± | 0.8 |
| 7500 ppm | 34 | 16.7± | 3.5 | 0.6± | 0.1 | 142± | 2 | 4.1± | 0.4* | 107± | 3 | 10.9± | 0.3 | 3.8± | 0.9 |
| 15000 ppm | 31 | 26.2± | 51.6 | 0.8± | 1.3 | 143± | 4 | 3.9± | 0.8 | 107± | 2 | 10.9± | 0.6 | 4.6± | 5.0 |
| 30000 ppm | 11 | 18.4± | 2.0* | 0.5± | 0.0 | 143± | 2 | 3.9± | 0.4 | 107± | 2 | 10.6± | 0.3 | 3.7± | 0.8 |

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

APPENDIX G 3

BIOCHEMISTRY (TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
SAMPLING DATE : 105-2
SEX : MALE

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of Animals | TOTAL PROTEIN g/dl | | ALBUMIN g/dl | | A/G RATIO | | T-BILIRUBIN mg/dl | | GLUCOSE mg/dl | | T-CHOLESTEROL mg/dl | | TRIGLYCERIDE mg/dl | |
|------------|-------------------|-----------------------|-----|-----------------|-----|-----------|-----|----------------------|------|------------------|----|------------------------|-----|-----------------------|----|
| Control | 31 | 5.6± | 0.8 | 3.0± | 0.5 | 1.2± | 0.2 | 0.18± | 0.05 | 176± | 35 | 125± | 61 | 39± | 13 |
| 10000 ppm | 35 | 5.4± | 0.6 | 3.0± | 0.4 | 1.2± | 0.2 | 0.18± | 0.07 | 174± | 33 | 110± | 25 | 47± | 21 |
| 20000 ppm | 27 | 5.5± | 0.8 | 3.0± | 0.4 | 1.2± | 0.2 | 0.22± | 0.16 | 156± | 42 | 158± | 173 | 44± | 21 |
| 40000 ppm | 25 | 5.2± | 0.6 | 2.8± | 0.4 | 1.2± | 0.2 | 0.18± | 0.05 | 162± | 34 | 120± | 41 | 46± | 18 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 SAMPLING DATE : 105-2
 SEX : MALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
 ALL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of Animals | GOT IU/ℓ | | GPT IU/ℓ | | LDH IU/ℓ | | ALP IU/ℓ | | CPK IU/ℓ | | UREA NITROGEN mg/dℓ | | SODIUM mEq/ℓ | |
|------------|-------------------|-------------|------|-------------|------|-------------|------|-------------|-----|-------------|-----|------------------------|-------|-----------------|---|
| Control | 31 | 128± | 194 | 84± | 144 | 488± | 736 | 180± | 118 | 59± | 74 | 26.9± | 15.0 | 156± | 2 |
| 10000 ppm | 35 | 214± | 630 | 94± | 203 | 493± | 778 | 175± | 83 | 46± | 23 | 20.7± | 2.4 | 156± | 2 |
| 20000 ppm | 27 | 733± | 1891 | 589± | 1489 | 2407± | 6369 | 309± | 432 | 59± | 57 | 20.7± | 2.9* | 157± | 2 |
| 40000 ppm | 25 | 182± | 278 | 128± | 177 | 690± | 947 | 238± | 373 | 68± | 73* | 21.5± | 8.5** | 157± | 2 |

Significant defference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
SAMPLING DATE : 105-2
SEX : MALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | POTASSIUM mEq/ℓ | | CHLORIDE mEq/ℓ | | CALCIUM mg/dℓ | | INORGANIC PHOSPHORUS mg/dℓ | |
|------------|-------------------|--------------------|-----|-------------------|---|------------------|-----|-------------------------------|-----|
| Control | 31 | 4.1± | 0.4 | 123± | 4 | 9.3± | 0.6 | 6.2± | 0.7 |
| 10000 ppm | 35 | 4.1± | 0.3 | 123± | 3 | 9.1± | 0.3 | 6.5± | 0.8 |
| 20000 ppm | 27 | 4.0± | 0.3 | 123± | 3 | 9.4± | 0.7 | 6.4± | 0.7 |
| 40000 ppm | 25 | 4.1± | 0.4 | 123± | 3 | 9.2± | 0.5 | 6.3± | 1.1 |

Significant defference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

APPENDIX G 4

BIOCHEMISTRY (TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
SAMPLING DATE : 105-2
SEX : FEMALE

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (105W)

REPORT TYPE : A1

PAGE : 4

| Group Name | NO. of Animals | TOTAL PROTEIN g / dl | | ALBUMIN g / dl | | A/G RATIO | | T-BILIRUBIN mg / dl | | GLUCOSE mg / dl | | T-CHOLESTEROL mg / dl | | TRIGLYCERIDE mg / dl | |
|------------|-------------------|-------------------------|-----|-------------------|-----|-----------|-----|------------------------|------|--------------------|----|--------------------------|----|-------------------------|----|
| Control | 28 | 5.3± | 0.7 | 2.9± | 0.4 | 1.2± | 0.2 | 0.17± | 0.02 | 133± | 21 | 92± | 47 | 46± | 27 |
| 10000 ppm | 24 | 5.2± | 0.8 | 2.9± | 0.1 | 1.3± | 0.2 | 0.18± | 0.03 | 137± | 31 | 73± | 12 | 49± | 61 |
| 20000 ppm | 18 | 5.4± | 1.0 | 2.9± | 0.3 | 1.2± | 0.3 | 0.17± | 0.03 | 130± | 30 | 92± | 86 | 48± | 31 |
| 40000 ppm | 26 | 5.6± | 1.3 | 2.9± | 0.3 | 1.2± | 0.3 | 0.17± | 0.02 | 129± | 24 | 78± | 34 | 45± | 28 |

Significant defference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
SAMPLING DATE : 105-2
SEX : FEMALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (105W)

PAGE : 5

| Group Name | NO. of Animals | GOT IU/ℓ | | GPT IU/ℓ | | LDH IU/ℓ | | ALP IU/ℓ | | CPK IU/ℓ | | UREA NITROGEN mg/dℓ | | SODIUM mEq/ℓ | |
|------------|-------------------|-------------|-----|-------------|----|-------------|------|-------------|-----|-------------|----|------------------------|-------|-----------------|---|
| Control | 28 | 91± | 25 | 38± | 14 | 301± | 226 | 208± | 114 | 64± | 43 | 15.3± | 2.3 | 154± | 2 |
| 10000 ppm | 24 | 88± | 35 | 35± | 16 | 265± | 70 | 235± | 79 | 79± | 88 | 16.8± | 3.3 | 154± | 2 |
| 20000 ppm | 18 | 141± | 206 | 39± | 21 | 1542± | 5305 | 194± | 70 | 80± | 69 | 18.1± | 3.5* | 155± | 2 |
| 40000 ppm | 26 | 96± | 38 | 47± | 32 | 347± | 203 | 214± | 89 | 65± | 45 | 21.1± | 7.7** | 155± | 2 |

Significant defference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
SAMPLING DATE : 105-2
SEX : FEMALE

REPORT TYPE : A1

BIOCHEMISTRY (SUMMARY)
ALL ANIMALS (105W)

PAGE : 6

| Group Name | NO. of Animals | POTASSIUM mEq/ℓ | | CHLORIDE mEq/ℓ | | CALCIUM mg/dℓ | | INORGANIC PHOSPHORUS mg/dℓ | |
|------------|-------------------|--------------------|------|-------------------|---|------------------|-----|-------------------------------|-----|
| Control | 28 | 4.1± | 0.3 | 124± | 4 | 9.4± | 0.6 | 6.7± | 1.0 |
| 10000 ppm | 24 | 4.1± | 0.3 | 124± | 2 | 9.2± | 0.4 | 6.5± | 0.8 |
| 20000 ppm | 18 | 4.0± | 0.3 | 123± | 3 | 9.4± | 0.7 | 6.7± | 1.0 |
| 40000 ppm | 26 | 3.9± | 0.3* | 123± | 3 | 9.4± | 0.6 | 6.7± | 0.9 |

Significant difference ; * : $P \leq 0.05$

** : $P \leq 0.01$

Test of Dunnett

(HCL074)

BAIS3

APPENDIX H 1

URINALYSIS (TOW-YEAR STUDY: SUMMARY)

RAT : MALE

STUDY NO. : 0224
 ANIMAL : RAT F344
 SAMPLING DATE : 104-1
 SEX : MALE

URINALYSIS

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of Animals | pH | | | | | | | CHI | Protein | | | | | CHI | Glucose | | | | | CHI | Ketone body | | | | | CHI | Bilirubin | | | | CHI | | | |
|------------|-------------------|-----|-----|-----|-----|-----|-----|-----|-----|---------|---|---|----|----|-----|---------|----|---|---|----|-----|-------------|----|---|----|---|-----|-----------|----|----|---|-----|---|----|----|
| | | 5.0 | 6.0 | 6.5 | 7.0 | 7.5 | 8.0 | 8.5 | | — | ± | + | 2+ | 3+ | | 4+ | — | ± | + | 2+ | | 3+ | 4+ | — | ± | + | | 2+ | 3+ | 4+ | — | | + | 2+ | 3+ |
| Control | 38 | 0 | 1 | 6 | 12 | 12 | 7 | 0 | | 0 | 0 | 0 | 1 | 18 | 19 | | 38 | 0 | 0 | 0 | 0 | 0 | 0 | | 36 | 2 | 0 | 0 | 0 | 0 | | 37 | 1 | 0 | 0 |
| 7500 ppm | 40 | 0 | 1 | 5 | 16 | 11 | 7 | 0 | | 0 | 0 | 1 | 0 | 6 | 33 | ** | 40 | 0 | 0 | 0 | 0 | 0 | 0 | | 40 | 0 | 0 | 0 | 0 | 0 | | 39 | 0 | 0 | 1 |
| 15000 ppm | 35 | 0 | 0 | 5 | 20 | 9 | 1 | 0 | | 0 | 0 | 0 | 0 | 0 | 35 | ** | 35 | 0 | 0 | 0 | 0 | 0 | 0 | | 35 | 0 | 0 | 0 | 0 | 0 | | 35 | 0 | 0 | 0 |
| 30000 ppm | 30 | 0 | 0 | 10 | 18 | 2 | 0 | 0 | ** | 0 | 0 | 0 | 0 | 1 | 29 | ** | 30 | 0 | 0 | 0 | 0 | 0 | 0 | | 30 | 0 | 0 | 0 | 0 | 0 | | 29 | 1 | 0 | 0 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

STUDY NO. : 0224
ANIMAL : RAT F344
SAMPLING DATE : 104-1
SEX : MALE

URINALYSIS

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of Animals | Occult blood | | | | | CHI | Urobilinogen | | | | | CHI |
|------------|-------------------|--------------|---|---|----|----|-----|--------------|---|----|----|----|-----|
| | | - | ± | + | 2+ | 3+ | | ± | + | 2+ | 3+ | 4+ | |
| Control | 38 | 38 | 0 | 0 | 0 | 0 | 0 | 38 | 0 | 0 | 0 | 0 | 0 |
| 7500 ppm | 40 | 39 | 1 | 0 | 0 | 0 | 0 | 40 | 0 | 0 | 0 | 0 | 0 |
| 15000 ppm | 35 | 33 | 1 | 0 | 0 | 0 | 1 | 35 | 0 | 0 | 0 | 0 | 0 |
| 30000 ppm | 30 | 30 | 0 | 0 | 0 | 0 | 0 | 30 | 0 | 0 | 0 | 0 | 0 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

APPENDIX H 2

URINALYSIS (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE

STUDY NO. : 0224

ANIMAL : RAT F344

SAMPLING DATE : 104-1

SEX : FEMALE

REPORT TYPE : A1

URINALYSIS

PAGE : 3

| Group Name | NO. of Animals | pH_____ | | | | | | | CHI | Protein_____ | | | | | CHI | Glucose_____ | | | | | CHI | Ketone body_____ | | | | | CHI | Bilirubin_____ | | | | CHI | | | |
|------------|-------------------|---------|-----|-----|-----|-----|-----|-----|-----|--------------|---|---|----|----|-----|--------------|----|---|---|----|-----|------------------|----|---|----|----|-----|----------------|----|----|----|-----|---|----|----|
| | | 5.0 | 6.0 | 6.5 | 7.0 | 7.5 | 8.0 | 8.5 | | — | ± | + | 2+ | 3+ | | 4+ | — | ± | + | 2+ | | 3+ | 4+ | — | ± | + | | 2+ | 3+ | 4+ | — | | + | 2+ | 3+ |
| Control | 47 | 0 | 1 | 11 | 13 | 15 | 5 | 2 | | 0 | 0 | 0 | 5 | 21 | 21 | | 47 | 0 | 0 | 0 | 0 | 0 | 0 | | 33 | 12 | 1 | 0 | 1 | 0 | | 46 | 0 | 0 | 1 |
| 7500 ppm | 39 | 0 | 0 | 3 | 17 | 16 | 3 | 0 | | 0 | 0 | 0 | 5 | 20 | 14 | | 39 | 0 | 0 | 0 | 0 | 0 | 0 | | 36 | 3 | 0 | 0 | 0 | 0 | | 39 | 0 | 0 | 0 |
| 15000 ppm | 35 | 0 | 0 | 7 | 16 | 9 | 2 | 1 | | 0 | 0 | 0 | 0 | 16 | 19 | | 35 | 0 | 0 | 0 | 0 | 0 | 0 | | 35 | 0 | 0 | 0 | 0 | 0 | ** | 34 | 0 | 0 | 1 |
| 30000 ppm | 14 | 0 | 0 | 5 | 4 | 4 | 1 | 0 | | 0 | 0 | 0 | 0 | 5 | 9 | | 14 | 0 | 0 | 0 | 0 | 0 | 0 | | 14 | 0 | 0 | 0 | 0 | 0 | | 14 | 0 | 0 | 0 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

STUDY NO. : 0224
ANIMAL : RAT F344
SAMPLING DATE : 104-1
SEX : FEMALE

URINALYSIS

REPORT TYPE : A1

PAGE : 4

| Group Name | NO. of Animals | Occult blood | | | | | Urabilinogen | | | | | | |
|------------|-------------------|--------------|---|---|----|----|--------------|----|---|----|----|----|-----|
| | | - | ± | + | 2+ | 3+ | CHI | ± | + | 2+ | 3+ | 4+ | CHI |
| Control | 47 | 42 | 4 | 0 | 1 | 0 | | 47 | 0 | 0 | 0 | 0 | |
| 7500 ppm | 39 | 38 | 0 | 0 | 1 | 0 | | 39 | 0 | 0 | 0 | 0 | |
| 15000 ppm | 35 | 35 | 0 | 0 | 0 | 0 | | 35 | 0 | 0 | 0 | 0 | |
| 30000 ppm | 14 | 14 | 0 | 0 | 0 | 0 | | 14 | 0 | 0 | 0 | 0 | |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

APPENDIX H 3

URINALYSIS (TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 SAMPLING DATE : 104-3
 SEX : MALE

URINALYSIS

REPORT TYPE : A1

PAGE : 1

| Group Name | NO. of Animals | pH | | | | | | | CHI | Protein | | | | | CHI | Glucose | | | | | CHI | Ketone body | | | | | CHI | Occult blood | | | | CHI | | | |
|------------|-------------------|-----|-----|-----|-----|-----|-----|-----|-----|---------|---|----|----|----|-----|---------|----|---|---|----|-----|-------------|----|----|----|---|-----|--------------|----|----|----|-----|---|---|----|
| | | 5.0 | 6.0 | 6.5 | 7.0 | 7.5 | 8.0 | 8.5 | | — | ± | + | 2+ | 3+ | | 4+ | — | ± | + | 2+ | | 3+ | 4+ | — | ± | + | | 2+ | 3+ | 4+ | — | | ± | + | 2+ |
| Control | 35 | 0 | 5 | 11 | 9 | 6 | 4 | 0 | | 0 | 3 | 21 | 10 | 1 | 0 | | 35 | 0 | 0 | 0 | 0 | 0 | | 14 | 21 | 0 | 0 | 0 | 0 | | 33 | 0 | 0 | 0 | 2 |
| 10000 ppm | 35 | 0 | 0 | 5 | 20 | 8 | 2 | 0 | * | 0 | 0 | 8 | 26 | 1 | 0 | ** | 35 | 0 | 0 | 0 | 0 | 0 | | 25 | 10 | 0 | 0 | 0 | 0 | ** | 34 | 0 | 0 | 0 | 1 |
| 20000 ppm | 30 | 0 | 0 | 9 | 14 | 3 | 4 | 0 | | 0 | 0 | 3 | 21 | 6 | 0 | ** | 30 | 0 | 0 | 0 | 0 | 0 | | 27 | 3 | 0 | 0 | 0 | 0 | ** | 29 | 1 | 0 | 0 | 0 |
| 40000 ppm | 25 | 0 | 0 | 5 | 19 | 1 | 0 | 0 | ** | 0 | 0 | 0 | 4 | 17 | 4 | ** | 25 | 0 | 0 | 0 | 0 | 0 | | 25 | 0 | 0 | 0 | 0 | 0 | ** | 24 | 1 | 0 | 0 | 0 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

STUDY NO. : 0225

URINALYSIS

ANIMAL : MOUSE BDF1

SAMPLING DATE : 104-3

SEX : MALE

REPORT TYPE : A1

PAGE : 2

| Group Name | NO. of Animals | Urobilinogen ± + 2+ 3+ 4+ CHI |
|------------|-------------------|----------------------------------|
| Control | 35 | 35 0 0 0 0 |
| 10000 ppm | 35 | 35 0 0 0 0 |
| 20000 ppm | 30 | 30 0 0 0 0 |
| 40000 ppm | 25 | 25 0 0 0 0 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

APPENDIX H 4

URINALYSIS (TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 SAMPLING DATE : 104-3
 SEX : FEMALE

URINALYSIS

REPORT TYPE : A1

PAGE : 3

| Group Name | NO. of Animals | pH_____ | | | | | | | CHI | Protein_____ | | | | | | CHI | Glucose_____ | | | | | | CHI | Ketone body_____ | | | | | | CHI | Occult blood_____ | | | | | | CHI |
|------------|-------------------|---------|-----|-----|-----|-----|-----|-----|-----|--------------|---|----|----|----|----|-----|--------------|---|---|----|----|----|-----|------------------|----|----|----|----|----|-----|-------------------|---|---|----|----|----|-----|
| | | 5.0 | 6.0 | 6.5 | 7.0 | 7.5 | 8.0 | 8.5 | | — | ± | + | 2+ | 3+ | 4+ | | — | ± | + | 2+ | 3+ | 4+ | | — | ± | + | 2+ | 3+ | 4+ | | — | ± | + | 2+ | 3+ | 4+ | |
| Control | 29 | 0 | 1 | 1 | 6 | 9 | 11 | 1 | | 0 | 0 | 20 | 6 | 2 | 1 | | 29 | 0 | 0 | 0 | 0 | 0 | | 1 | 11 | 16 | 1 | 0 | 0 | | 25 | 1 | 2 | 1 | 0 | | |
| 10000 ppm | 25 | 0 | 0 | 1 | 8 | 8 | 8 | 0 | | 0 | 1 | 3 | 14 | 6 | 1 | ** | 25 | 0 | 0 | 0 | 0 | 0 | | 16 | 9 | 0 | 0 | 0 | 0 | ** | 18 | 4 | 1 | 1 | 1 | | |
| 20000 ppm | 19 | 0 | 0 | 0 | 6 | 8 | 5 | 0 | | 0 | 0 | 2 | 6 | 7 | 4 | ** | 19 | 0 | 0 | 0 | 0 | 0 | | 17 | 2 | 0 | 0 | 0 | 0 | ** | 15 | 3 | 0 | 0 | 1 | | |
| 40000 ppm | 31 | 0 | 0 | 5 | 9 | 13 | 4 | 0 | | 0 | 0 | 0 | 2 | 6 | 23 | ** | 31 | 0 | 0 | 0 | 0 | 0 | | 31 | 0 | 0 | 0 | 0 | 0 | ** | 21 | 1 | 2 | 4 | 3 | | |

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

STUDY NO. : 0225

ANIMAL : MOUSE BDF1

SAMPLING DATE : 104-3

SEX : FEMALE

REPORT TYPE : A1

URINALYSIS

PAGE : 4

| Group Name | NO. of Animals | Urobilinogen ± + 2+ 3+ 4+ CHI |
|------------|-------------------|----------------------------------|
| Control | 29 | 29 0 0 0 0 |
| 10000 ppm | 25 | 25 0 0 0 0 |
| 20000 ppm | 19 | 19 0 0 0 0 |
| 40000 ppm | 31 | 31 0 0 0 0 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of CHI SQUARE

(HCL101)

BAIS3

APPENDIX I 1

GROSS FINDINGS (TOW-YEAR STUDY: SUMMARY)

RAT : MALE : DEAD AND MORIBUND ANIMALS

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 1

| Organ | Findings | Group Name NO. of Animals | Control 12 (%) | 7500 ppm 11 (%) | 15000 ppm 16 (%) | 30000 ppm 20 (%) |
|-------------|-------------|------------------------------|-------------------|--------------------|---------------------|---------------------|
| skin/app | nodule | | 1 (8) | 0 (0) | 0 (0) | 2 (10) |
| subcutis | jaundice | | 1 (8) | 0 (0) | 1 (6) | 1 (5) |
| | mass | | 1 (8) | 4 (36) | 3 (19) | 5 (25) |
| lung | red | | 0 (0) | 1 (9) | 1 (6) | 0 (0) |
| | white zone | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |
| | red zone | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| lymph node | enlarged | | 1 (8) | 0 (0) | 1 (6) | 1 (5) |
| spleen | enlarged | | 5 (42) | 0 (0) | 4 (25) | 2 (10) |
| | white zone | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | nodule | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| heart | white zone | | 0 (0) | 0 (0) | 0 (0) | 5 (25) |
| | dilated | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| forestomach | ulcer | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| st stomach | red zone | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | ulcer | | 0 (0) | 1 (9) | 3 (19) | 0 (0) |
| liver | enlarged | | 3 (25) | 0 (0) | 2 (13) | 1 (5) |
| | yellow | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | yellow zone | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| | nodule | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| | rough | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| pancreas | nodule | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 2

| Organ | Findings | Group Name NO. of Animals | Control 12 (%) | 7500 ppm 11 (%) | 15000 ppm 16 (%) | 30000 ppm 20 (%) |
|-------------|---------------|------------------------------|-------------------|--------------------|---------------------|---------------------|
| kidney | pale | | 0 (0) | 0 (0) | 0 (0) | 2 (10) |
| | white zone | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| | granular | | 2 (17) | 1 (9) | 2 (13) | 3 (15) |
| urin bladd | urine:red | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| pituitary | enlarged | | 2 (17) | 2 (18) | 3 (19) | 2 (10) |
| | red zone | | 1 (8) | 0 (0) | 0 (0) | 3 (15) |
| | nodule | | 0 (0) | 1 (9) | 0 (0) | 3 (15) |
| adrenal | enlarged | | 1 (8) | 0 (0) | 1 (6) | 1 (5) |
| testis | atrophic | | 1 (8) | 0 (0) | 2 (13) | 1 (5) |
| | nodule | | 8 (67) | 3 (27) | 8 (50) | 9 (45) |
| prostate | brown | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| brain | red zone | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | elevated | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |
| spinal cord | red zone | | 1 (8) | 0 (0) | 0 (0) | 0 (0) |
| | hemorrhage | | 0 (0) | 0 (0) | 1 (6) | 0 (0) |
| eye | white | | 1 (8) | 0 (0) | 1 (6) | 2 (10) |
| Zymbal gl | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| muscle | nodule | | 1 (8) | 0 (0) | 1 (6) | 0 (0) |
| peritoneum | nodule | | 0 (0) | 3 (27) | 4 (25) | 2 (10) |
| abdominal c | brown | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| | ascites | | 0 (0) | 1 (9) | 2 (13) | 0 (0) |
| thoracic ca | pleural fluid | | 1 (8) | 1 (9) | 1 (6) | 1 (5) |

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 3

| Organ | Findings | Group Name | Control | 7500 ppm | 15000 ppm | 30000 ppm |
|------------|-----------------|----------------|---------|----------|-----------|-----------|
| | | NO. of Animals | 12 (%) | 11 (%) | 16 (%) | 20 (%) |
| other | hindlimb:nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| whole body | anemic | | 1 (8) | 0 (0) | 1 (6) | 0 (0) |

(HPT080)

BAIS 3

APPENDIX I 2

GROSS FINDINGS (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE : DEAD AND MORIBUND ANIMALS

STUDY NO. : 0224
 ANIMAL : RAT F344
 REPORT TYPE : A1
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 4

| Organ | Findings | Group Name NO. of Animals | Control 3 (%) | 7500 ppm 11 (%) | 15000 ppm 15 (%) | 30000 ppm 36 (%) |
|-------------|------------|------------------------------|------------------|--------------------|---------------------|---------------------|
| subcutis | jaundice | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| | dry | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| | mass | | 1 (33) | 4 (36) | 3 (20) | 4 (11) |
| lung | white zone | | 0 (0) | 0 (0) | 1 (7) | 1 (3) |
| | red zone | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |
| | voluminous | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| lymph node | enlarged | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |
| spleen | enlarged | | 0 (0) | 4 (36) | 1 (7) | 3 (8) |
| | atrophic | | 0 (0) | 0 (0) | 0 (0) | 2 (6) |
| | adhesion | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| heart | white zone | | 0 (0) | 0 (0) | 0 (0) | 6 (17) |
| | dilated | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| oral cavity | ulcer | | 1 (33) | 0 (0) | 0 (0) | 0 (0) |
| forestomach | ulcer | | 0 (0) | 1 (9) | 1 (7) | 1 (3) |
| | thick | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |
| gl stomach | ulcer | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| large intes | nodule | | 0 (0) | 0 (0) | 1 (7) | 0 (0) |
| liver | white zone | | 0 (0) | 0 (0) | 1 (7) | 0 (0) |
| | nodule | | 1 (33) | 0 (0) | 0 (0) | 1 (3) |
| | rough | | 0 (0) | 3 (27) | 0 (0) | 1 (3) |
| | granular | | 0 (0) | 0 (0) | 1 (7) | 0 (0) |
| | herniation | | 0 (0) | 0 (0) | 0 (0) | 2 (6) |

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 5

| Organ | Findings | Group Name NO. of Animals | Control 3 (%) | 7500 ppm 11 (%) | 15000 ppm 15 (%) | 30000 ppm 36 (%) |
|-------------|---------------|------------------------------|------------------|--------------------|---------------------|---------------------|
| kidney | turbid | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| | pale | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| | deformed | | 0 (0) | 0 (0) | 1 (7) | 0 (0) |
| | granular | | 1 (33) | 1 (9) | 1 (7) | 0 (0) |
| pituitary | enlarged | | 1 (33) | 4 (36) | 2 (13) | 2 (6) |
| | red zone | | 0 (0) | 0 (0) | 1 (7) | 2 (6) |
| | nodule | | 0 (0) | 0 (0) | 1 (7) | 3 (8) |
| thyroid | enlarged | | 0 (0) | 0 (0) | 1 (7) | 0 (0) |
| adrenal | enlarged | | 1 (33) | 0 (0) | 0 (0) | 0 (0) |
| ovary | cyst | | 0 (0) | 0 (0) | 1 (7) | 1 (3) |
| uterus | nodule | | 0 (0) | 0 (0) | 6 (40) | 2 (6) |
| | dilated lumen | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| vagina | nodule | | 0 (0) | 0 (0) | 1 (7) | 2 (6) |
| prep/cli gl | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| spinal cord | red zone | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |
| eye | white | | 1 (33) | 1 (9) | 0 (0) | 2 (6) |
| | red | | 0 (0) | 0 (0) | 1 (7) | 0 (0) |
| Harder gl | enlarged | | 1 (33) | 0 (0) | 0 (0) | 0 (0) |
| Zymbal gl | nodule | | 0 (0) | 0 (0) | 1 (7) | 0 (0) |
| abdominal c | hemorrhage | | 0 (0) | 1 (9) | 0 (0) | 0 (0) |
| | mass | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| | ascites | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 6

| Organ_____ | Findings_____ | Group Name NO. of Animals | Control 3 (%) | 7500 ppm 11 (%) | 15000 ppm 15 (%) | 30000 ppm 36 (%) |
|-------------|---------------|------------------------------|------------------|--------------------|---------------------|---------------------|
| adipose | white zone | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| thoracic ca | pleural fluid | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| whole body | anemic | | 0 (0) | 1 (9) | 1 (7) | 0 (0) |

(HPT080)

BAIS 3

APPENDIX I 3

GROSS FINDINGS (TOW-YEAR STUDY: SUMMARY)

RAT : MALE : SACRIFICED ANIMALS

STUDY NO. : 0224
 ANIMAL : RAT F344
 REPORT TYPE : A1
 SEX : MALE

GROSS FINDINGS (SUMMARY)
 SACRIFICED ANIMALS (105W)

PAGE : 1

| Organ | Findings | Group Name NO. of Animals | Control 38 (%) | 7500 ppm 39 (%) | 15000 ppm 34 (%) | 30000 ppm 30 (%) |
|-------------|------------|------------------------------|-------------------|--------------------|---------------------|---------------------|
| skin/app | nodule | | 2 (5) | 6 (15) | 1 (3) | 7 (23) |
| | scab | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| subcutis | mass | | 8 (21) | 6 (15) | 5 (15) | 3 (10) |
| nasal cavit | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| lung | nodule | | 2 (5) | 1 (3) | 5 (15) | 5 (17) |
| lymph node | enlarged | | 0 (0) | 1 (3) | 0 (0) | 1 (3) |
| spleen | enlarged | | 1 (3) | 2 (5) | 0 (0) | 3 (10) |
| | deformed | | 0 (0) | 2 (5) | 1 (3) | 0 (0) |
| salivary gl | cyst | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| forestomach | nodule | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | ulcer | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| small intes | nodule | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| large intes | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| liver | white zone | | 0 (0) | 1 (3) | 0 (0) | 1 (3) |
| | nodule | | 1 (3) | 1 (3) | 0 (0) | 2 (7) |
| | rough | | 0 (0) | 2 (5) | 0 (0) | 1 (3) |
| | herniation | | 0 (0) | 1 (3) | 1 (3) | 1 (3) |
| pancreas | nodule | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| kidney | cyst | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| | granular | | 20 (53) | 22 (56) | 15 (44) | 2 (7) |
| pituitary | enlarged | | 3 (8) | 2 (5) | 1 (3) | 0 (0) |
| | red zone | | 0 (0) | 0 (0) | 1 (3) | 1 (3) |

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 2

| Organ | Findings | Group Name NO. of Animals | Control 38 (%) | 7500 ppm 39 (%) | 15000 ppm 34 (%) | 30000 ppm 30 (%) |
|-------------|------------|------------------------------|-------------------|--------------------|---------------------|---------------------|
| pituitary | nodule | | 5 (13) | 4 (10) | 3 (9) | 3 (10) |
| thyroid | enlarged | | 1 (3) | 2 (5) | 4 (12) | 0 (0) |
| | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| adrenal | enlarged | | 2 (5) | 0 (0) | 1 (3) | 1 (3) |
| testis | nodule | | 35 (92) | 37 (95) | 34 (100) | 27 (90) |
| eye | white | | 3 (8) | 5 (13) | 5 (15) | 0 (0) |
| | red | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| vertebra | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (3) |
| peritoneum | nodule | | 1 (3) | 2 (5) | 0 (0) | 1 (3) |
| retroperit | edema | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| abdominal c | ascites | | 0 (0) | 2 (5) | 1 (3) | 1 (3) |
| other | lip:nodule | | 1 (3) | 0 (0) | 0 (0) | 1 (3) |
| | ear:nodule | | 0 (0) | 1 (3) | 1 (3) | 0 (0) |

APPENDIX I 4

GROSS FINDINGS (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE : SACRIFICED ANIMALS

STUDY NO. : 0224
 ANIMAL : RAT F344
 REPORT TYPE : A1
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)
 SACRIFICED ANIMALS (105W)

PAGE : 3

| Organ | Findings | Group Name NO. of Animals | Control 47 (%) | 7500 ppm 39 (%) | 15000 ppm 35 (%) | 30000 ppm 14 (%) |
|-------------|-------------|------------------------------|-------------------|--------------------|---------------------|---------------------|
| skin/app | nodule | | 2 (4) | 0 (0) | 0 (0) | 1 (7) |
| | scab | | 1 (2) | 1 (3) | 0 (0) | 0 (0) |
| subcutis | mass | | 7 (15) | 7 (18) | 3 (9) | 2 (14) |
| lung | nodule | | 1 (2) | 2 (5) | 0 (0) | 0 (0) |
| lymph node | enlarged | | 0 (0) | 2 (5) | 0 (0) | 0 (0) |
| spleen | enlarged | | 1 (2) | 3 (8) | 1 (3) | 1 (7) |
| | deformed | | 2 (4) | 0 (0) | 0 (0) | 0 (0) |
| heart | white zone | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| oral cavity | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| small intes | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| liver | white patch | | 1 (2) | 0 (0) | 0 (0) | 0 (0) |
| | nodule | | 1 (2) | 0 (0) | 0 (0) | 0 (0) |
| | deformed | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | rough | | 2 (4) | 2 (5) | 1 (3) | 1 (7) |
| | herniation | | 2 (4) | 3 (8) | 1 (3) | 0 (0) |
| pancreas | nodule | | 1 (2) | 0 (0) | 0 (0) | 0 (0) |
| kidney | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (7) |
| | granular | | 4 (9) | 5 (13) | 3 (9) | 0 (0) |
| pituitary | enlarged | | 3 (6) | 3 (8) | 1 (3) | 0 (0) |
| | red zone | | 3 (6) | 3 (8) | 5 (14) | 1 (7) |
| | nodule | | 6 (13) | 3 (8) | 3 (9) | 1 (7) |
| thyroid | enlarged | | 1 (2) | 1 (3) | 1 (3) | 0 (0) |

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 4

| Organ | Findings | Group Name NO. of Animals | Control 47 (%) | 7500 ppm 39 (%) | 15000 ppm 35 (%) | 30000 ppm 14 (%) |
|-------------|------------|------------------------------|-------------------|--------------------|---------------------|---------------------|
| adrenal | enlarged | | 0 (0) | 2 (5) | 0 (0) | 0 (0) |
| ovary | enlarged | | 0 (0) | 1 (3) | 2 (6) | 0 (0) |
| | nodule | | 1 (2) | 0 (0) | 0 (0) | 0 (0) |
| | cyst | | 1 (2) | 2 (5) | 0 (0) | 0 (0) |
| uterus | nodule | | 10 (21) | 4 (10) | 3 (9) | 2 (14) |
| eye | white | | 1 (2) | 1 (3) | 3 (9) | 0 (0) |
| Zymbal gl | nodule | | 1 (2) | 0 (0) | 0 (0) | 0 (0) |
| thoracic ca | hemorrhage | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| other | lip:nodule | | 0 (0) | 1 (3) | 2 (6) | 1 (7) |
| | ear:nodule | | 1 (2) | 0 (0) | 0 (0) | 0 (0) |

(HPT080)

BAIS3

APPENDIX I 5

GROSS FINDINGS(TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE : DEAD AND MORIBUND ANIMALS

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 REPORT TYPE : A1
 SEX : MALE

GROSS FINDINGS (SUMMARY)
 DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 1

| Organ | Findings | Group Name NO. of Animals | Control 16 (%) | 10000 ppm 15 (%) | 20000 ppm 20 (%) | 40000 ppm 25 (%) |
|-------------|------------|------------------------------|-------------------|---------------------|---------------------|---------------------|
| skin/app | ulcer | | 0 (0) | 2 (13) | 1 (5) | 3 (12) |
| subcutis | edema | | 0 (0) | 1 (7) | 0 (0) | 2 (8) |
| | mass | | 1 (6) | 3 (20) | 2 (10) | 1 (4) |
| | thick | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| lung | red zone | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| | nodule | | 2 (13) | 3 (20) | 3 (15) | 1 (4) |
| lymph node | enlarged | | 3 (19) | 0 (0) | 2 (10) | 4 (16) |
| spleen | enlarged | | 2 (13) | 0 (0) | 0 (0) | 5 (20) |
| | red zone | | 0 (0) | 0 (0) | 1 (5) | 0 (0) |
| | black zone | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | nodule | | 0 (0) | 2 (13) | 1 (5) | 0 (0) |
| | deformed | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | adhesion | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| heart | white zone | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| salivary gl | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| gl stomach | white | | 1 (6) | 0 (0) | 0 (0) | 0 (0) |
| small intes | nodule | | 0 (0) | 0 (0) | 1 (5) | 0 (0) |
| | adhesion | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| large intes | adhesion | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| liver | enlarged | | 1 (6) | 0 (0) | 1 (5) | 3 (12) |
| | pale | | 1 (6) | 1 (7) | 0 (0) | 0 (0) |
| | white zone | | 0 (0) | 1 (7) | 3 (15) | 4 (16) |

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 2

| Organ | Findings | Group Name NO. of Animals | Control | 10000 ppm | 20000 ppm | 40000 ppm |
|-------------|------------------------|------------------------------|---------|-----------|-----------|-----------|
| | | | 16 (%) | 15 (%) | 20 (%) | 25 (%) |
| liver | red zone | | 0 (0) | 1 (7) | 1 (5) | 0 (0) |
| | nodule | | 5 (31) | 6 (40) | 7 (35) | 7 (28) |
| | deformed | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| | adhesion | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| pancreas | thick | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| kidney | atrophic | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | pale | | 1 (6) | 0 (0) | 1 (5) | 1 (4) |
| | white patch | | 1 (6) | 0 (0) | 0 (0) | 0 (0) |
| | hydronephrosis | | 2 (13) | 0 (0) | 0 (0) | 0 (0) |
| urin bladd | red zone | | 1 (6) | 0 (0) | 0 (0) | 0 (0) |
| | dilated | | 1 (6) | 0 (0) | 0 (0) | 0 (0) |
| | urine:marked retention | | 3 (19) | 4 (27) | 4 (20) | 3 (12) |
| testis | atrophic | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| epididymis | red | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | nodule | | 0 (0) | 1 (7) | 1 (5) | 0 (0) |
| | adhesion | | 0 (0) | 1 (7) | 0 (0) | 0 (0) |
| semin ves | red | | 1 (6) | 0 (0) | 0 (0) | 0 (0) |
| | adhesion | | 0 (0) | 1 (7) | 1 (5) | 0 (0) |
| prep/cli gl | enlarged | | 0 (0) | 1 (7) | 0 (0) | 1 (4) |
| | nodule | | 0 (0) | 0 (0) | 1 (5) | 0 (0) |
| brain | swollen | | 0 (0) | 0 (0) | 1 (5) | 0 (0) |
| | red zone | | 1 (6) | 1 (7) | 1 (5) | 0 (0) |

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 3

| Organ | Findings | Group Name NO. of Animals | Control | 10000 ppm | 20000 ppm | 40000 ppm |
|-------------|-----------------|------------------------------|---------|-----------|-----------|-----------|
| | | | 16 (%) | 15 (%) | 20 (%) | 25 (%) |
| periph nerv | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| Harder gl | enlarged | | 1 (6) | 0 (0) | 1 (5) | 0 (0) |
| | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| mediastinum | mass | | 0 (0) | 0 (0) | 0 (0) | 2 (8) |
| abdominal c | hemorrhage | | 0 (0) | 0 (0) | 2 (10) | 0 (0) |
| | ascites | | 1 (6) | 3 (20) | 0 (0) | 2 (8) |
| mesenterium | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| thoracic ca | hemorrhage | | 0 (0) | 1 (7) | 0 (0) | 1 (4) |
| | pleural fluid | | 3 (19) | 3 (20) | 0 (0) | 2 (8) |
| other | red zone | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | hindlimb:nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |

APPENDIX I 6

GROSS FINDINGS (TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE : DEAD AND MORIBUND ANIMALS

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 4

| Organ | Findings | Group Name NO. of Animals | Control | 10000 ppm | 20000 ppm | 40000 ppm |
|-------------|----------------------------|------------------------------|---------|-----------|-----------|-----------|
| | | | 21 (%) | 26 (%) | 31 (%) | 22 (%) |
| subcutis | red zone | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| | edema | | 5 (24) | 5 (19) | 6 (19) | 3 (14) |
| | nodule | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |
| | mass | | 2 (10) | 4 (15) | 3 (10) | 6 (27) |
| lung | red zone | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | edema | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| | nodule | | 1 (5) | 0 (0) | 1 (3) | 2 (9) |
| lymph node | enlarged | | 3 (14) | 8 (31) | 6 (19) | 3 (14) |
| | white | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| thymus | enlarged | | 1 (5) | 0 (0) | 1 (3) | 0 (0) |
| spleen | enlarged | | 6 (29) | 8 (31) | 7 (23) | 4 (18) |
| | nodule | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |
| | accentuation of white pulp | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |
| heart | dilated | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |
| forestomach | nodule | | 1 (5) | 0 (0) | 0 (0) | 1 (5) |
| small intes | nodule | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| liver | enlarged | | 4 (19) | 2 (8) | 9 (29) | 1 (5) |
| | white zone | | 4 (19) | 4 (15) | 8 (26) | 1 (5) |
| | red zone | | 2 (10) | 0 (0) | 1 (3) | 2 (9) |
| | nodule | | 1 (5) | 5 (19) | 1 (3) | 6 (27) |
| | rough | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| | nodular | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 5

| Organ | Findings | Group Name NO. of Animals | Control | 10000 ppm | 20000 ppm | 40000 ppm |
|-------------|------------------------|------------------------------|---------|-----------|-----------|-----------|
| | | | 21 (%) | 26 (%) | 31 (%) | 22 (%) |
| Liver | adhesion | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| pancreas | nodule | | 2 (10) | 0 (0) | 3 (10) | 0 (0) |
| kidney | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| | pale | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| | black zone | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| | nodule | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |
| | granular | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| | hydronephrosis | | 2 (10) | 0 (0) | 2 (6) | 1 (5) |
| ureter | dilated | | 0 (0) | 0 (0) | 0 (0) | 1 (5) |
| urin bladd | urine:marked retention | | 0 (0) | 1 (4) | 0 (0) | 1 (5) |
| pituitary | enlarged | | 2 (10) | 1 (4) | 2 (6) | 2 (9) |
| | nodule | | 0 (0) | 1 (4) | 2 (6) | 0 (0) |
| ovary | enlarged | | 6 (29) | 3 (12) | 5 (16) | 6 (27) |
| | cyst | | 1 (5) | 3 (12) | 2 (6) | 1 (5) |
| uterus | nodule | | 7 (33) | 3 (12) | 10 (32) | 8 (36) |
| brain | red zone | | 0 (0) | 1 (4) | 1 (3) | 0 (0) |
| periph nerv | nodule | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |
| Harder gl | nodule | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| mediastinum | nodule | | 1 (5) | 1 (4) | 0 (0) | 0 (0) |
| | mass | | 1 (5) | 1 (4) | 2 (6) | 3 (14) |
| peritoneum | nodule | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | mass | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
DEAD AND MORIBUND ANIMALS (0-105W)

PAGE : 6

| Organ | Findings | Group Name NO. of Animals | Control | 10000 ppm | 20000 ppm | 40000 ppm |
|-------------|-----------------|------------------------------|---------|-----------|-----------|-----------|
| | | | 21 (%) | 26 (%) | 31 (%) | 22 (%) |
| peritoneum | thick | | 1 (5) | 2 (8) | 1 (3) | 1 (5) |
| retroperit | mass | | 0 (0) | 1 (4) | 0 (0) | 1 (5) |
| abdominal c | hemorrhage | | 0 (0) | 1 (4) | 3 (10) | 1 (5) |
| | ascites | | 8 (38) | 7 (27) | 12 (39) | 7 (32) |
| thoracic ca | pleural fluid | | 6 (29) | 13 (50) | 13 (42) | 12 (55) |
| other | hindlimb:nodule | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |
| whole body | anemic | | 1 (5) | 0 (0) | 0 (0) | 0 (0) |

(HPT080)

BAIS 3

APPENDIX I 7

GROSS FINDINGS (TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE : SACRIFICED ANIMALS

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 1

| Organ | Findings | Group Name NO. of Animals | Control 34 (%) | 10000 ppm 35 (%) | 20000 ppm 30 (%) | 40000 ppm 25 (%) |
|-------------|------------------------|------------------------------|-------------------|---------------------|---------------------|---------------------|
| skin/app | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| | ulcer | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | erosion | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| | scab | | 1 (3) | 1 (3) | 0 (0) | 0 (0) |
| subcutis | mass | | 2 (6) | 0 (0) | 1 (3) | 0 (0) |
| lung | white zone | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | nodule | | 4 (12) | 5 (14) | 7 (23) | 5 (20) |
| lymph node | enlarged | | 2 (6) | 3 (9) | 2 (7) | 1 (4) |
| spleen | enlarged | | 2 (6) | 1 (3) | 2 (7) | 0 (0) |
| | black zone | | 1 (3) | 4 (11) | 1 (3) | 1 (4) |
| | nodule | | 1 (3) | 0 (0) | 1 (3) | 1 (4) |
| | deformed | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| tongue | nodule | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| forestomach | nodule | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | ulcer | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| liver | red zone | | 0 (0) | 1 (3) | 1 (3) | 0 (0) |
| | nodule | | 17 (50) | 21 (60) | 16 (53) | 14 (56) |
| pancreas | nodule | | 2 (6) | 0 (0) | 1 (3) | 1 (4) |
| kidney | atrophic | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | white zone | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| | hydronephrosis | | 1 (3) | 0 (0) | 1 (3) | 0 (0) |
| urin bladd | urine:marked retention | | 2 (6) | 0 (0) | 0 (0) | 0 (0) |

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 2

| Organ | Findings | Group Name NO. of Animals | Control | 10000 ppm | 20000 ppm | 40000 ppm |
|-------------|-----------------|------------------------------|---------|-----------|-----------|-----------|
| | | | 34 (%) | 35 (%) | 30 (%) | 25 (%) |
| adrenal | enlarged | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| testis | enlarged | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| epididymis | nodule | | 0 (0) | 0 (0) | 1 (3) | 0 (0) |
| prep/cli gl | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| Harder gl | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | nodule | | 0 (0) | 1 (3) | 1 (3) | 0 (0) |
| pleura | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| peritoneum | nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| other | tail:nodule | | 0 (0) | 1 (3) | 0 (0) | 0 (0) |
| | hindlimb:nodule | | 0 (0) | 0 (0) | 1 (3) | 1 (4) |

(HPT080)

BAIS3

APPENDIX I 8

GROSS FINDINGS(TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE : SACRIFICED ANIMALS

STUDY NO. : 0225
 ANIMAL : MOUSE BDF1
 REPORT TYPE : A1
 SEX : FEMALE

GROSS FINDINGS (SUMMARY)
 SACRIFICED ANIMALS (105W)

PAGE : 3

| Organ | Findings | Group Name NO. of Animals | Control 29 (%) | 10000 ppm 24 (%) | 20000 ppm 19 (%) | 40000 ppm 28 (%) |
|-------------|----------------|------------------------------|-------------------|---------------------|---------------------|---------------------|
| skin/app | nodule | | 1 (3) | 0 (0) | 0 (0) | 1 (4) |
| | scab | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| subcutis | mass | | 2 (7) | 0 (0) | 0 (0) | 1 (4) |
| lung | nodule | | 1 (3) | 3 (13) | 1 (5) | 2 (7) |
| lymph node | enlarged | | 4 (14) | 1 (4) | 3 (16) | 2 (7) |
| spleen | enlarged | | 1 (3) | 1 (4) | 1 (5) | 3 (11) |
| | white zone | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | black zone | | 1 (3) | 1 (4) | 0 (0) | 0 (0) |
| | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | deformed | | 1 (3) | 0 (0) | 1 (5) | 0 (0) |
| tongue | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| salivary gl | enlarged | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |
| forestomach | nodule | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |
| gl stomach | ulcer | | 0 (0) | 0 (0) | 1 (5) | 0 (0) |
| duodenum | nodule | | 0 (0) | 0 (0) | 1 (5) | 1 (4) |
| liver | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | red zone | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | nodule | | 4 (14) | 9 (38) | 6 (32) | 8 (29) |
| | rough | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| kidney | white zone | | 0 (0) | 1 (4) | 0 (0) | 1 (4) |
| | hydronephrosis | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| urin bladd | nodule | | 0 (0) | 1 (4) | 0 (0) | 0 (0) |

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE

GROSS FINDINGS (SUMMARY)
SACRIFICED ANIMALS (105W)

PAGE : 4

| Organ | Findings | Group Name NO. of Animals | Control | 10000 ppm | 20000 ppm | 40000 ppm |
|-------------|---------------|------------------------------|---------|-----------|-----------|-----------|
| | | | 29 (%) | 24 (%) | 19 (%) | 28 (%) |
| pituitary | enlarged | | 3 (10) | 2 (8) | 2 (11) | 0 (0) |
| | red zone | | 0 (0) | 0 (0) | 1 (5) | 0 (0) |
| | nodule | | 6 (21) | 3 (13) | 2 (11) | 6 (21) |
| ovary | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| | cyst | | 3 (10) | 4 (17) | 4 (21) | 2 (7) |
| uterus | nodule | | 2 (7) | 3 (13) | 3 (16) | 3 (11) |
| eye | white | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| Harder gl | enlarged | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| peritoneum | nodule | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| abdominal c | ascites | | 0 (0) | 4 (17) | 4 (21) | 2 (7) |
| thoracic ca | pleural fluid | | 0 (0) | 0 (0) | 0 (0) | 1 (4) |
| other | scab | | 1 (3) | 0 (0) | 0 (0) | 0 (0) |

(HPT080)

BAIS 3

APPENDIX J 1

ORGAN WEIGHT, ABSOLUTE, (TOW-YEAR STUDY: SUMMARY)

RAT : MALE

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of Animals | Body Weight | | ADRENALS | | TESTES | | HEART | | LUNGS | | KIDNEYS | |
|------------|-------------------|-------------|------|----------|-------|--------|-------|--------|--------|--------|-------|---------|---------|
| Control | 38 | 420± | 50 | 0.090± | 0.047 | 4.132± | 1.444 | 1.236± | 0.120 | 1.460± | 0.111 | 3.099± | 0.533 |
| 7500 ppm | 39 | 418± | 34 | 0.082± | 0.015 | 4.481± | 1.550 | 1.229± | 0.105 | 1.495± | 0.185 | 3.136± | 0.420 |
| 15000 ppm | 34 | 403± | 38 | 0.164± | 0.481 | 4.822± | 1.464 | 1.195± | 0.161 | 1.505± | 0.178 | 3.070± | 0.419 |
| 30000 ppm | 30 | 378± | 40** | 0.083± | 0.034 | 4.901± | 1.643 | 1.145± | 0.118* | 1.515± | 0.424 | 2.788± | 0.136** |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS3

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of Animals | SPLEEN | | LIVER | | BRAIN | |
|------------|-------------------|--------|-------|---------|---------|--------|-------|
| Control | 38 | 1.180± | 0.375 | 13.268± | 1.663 | 2.025± | 0.041 |
| 7500 ppm | 39 | 1.587± | 2.032 | 13.622± | 2.043 | 2.016± | 0.057 |
| 15000 ppm | 34 | 1.214± | 0.257 | 12.783± | 1.441 | 2.007± | 0.056 |
| 30000 ppm | 30 | 1.415± | 1.132 | 12.168± | 4.073** | 2.006± | 0.055 |

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS3

APPENDIX J 2

ORGAN WEIGHT, ABSOLUTE (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | Body Weight | | ADRENALS | | OVARIES | | HEART | | LUNGS | | KIDNEYS | |
|------------|-------------------|-------------|------|----------|-------|---------|-------|--------|-------|--------|-------|---------|-------|
| Control | 47 | 294± | 34 | 0.071± | 0.009 | 0.169± | 0.191 | 0.891± | 0.071 | 1.028± | 0.073 | 1.941± | 0.160 |
| 7500 ppm | 39 | 293± | 31 | 0.094± | 0.104 | 0.220± | 0.450 | 0.905± | 0.052 | 1.122± | 0.301 | 1.981± | 0.223 |
| 15000 ppm | 35 | 276± | 38* | 0.077± | 0.020 | 0.384± | 1.322 | 0.872± | 0.063 | 1.039± | 0.096 | 2.013± | 0.215 |
| 30000 ppm | 14 | 258± | 20** | 0.072± | 0.007 | 0.135± | 0.024 | 0.863± | 0.045 | 1.083± | 0.325 | 2.043± | 0.146 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS 3

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | SPLEEN | | LIVER | | BRAIN | |
|------------|-------------------|--------|-------|--------|-------|--------|-------|
| Control | 47 | 0.646± | 0.348 | 7.667± | 1.118 | 1.839± | 0.056 |
| 7500 ppm | 39 | 0.801± | 0.912 | 7.560± | 1.041 | 1.835± | 0.060 |
| 15000 ppm | 35 | 0.652± | 0.327 | 7.330± | 0.987 | 1.829± | 0.036 |
| 30000 ppm | 14 | 0.803± | 0.924 | 7.202± | 1.142 | 1.836± | 0.053 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

(HCL040)

BAIS3

APPENDIX J 3

ORGAN WEIGHT, ABSOLUTE (TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of Animals | Body Weight | ADRENALS | | TESTES | | HEART | | LUNGS | | KIDNEYS | |
|------------|-------------------|-------------|----------|-------|--------|-------|--------|---------|--------|-------|---------|-------|
| Control | 34 | 45.2± 8.5 | 0.013± | 0.009 | 0.224± | 0.073 | 0.224± | 0.046 | 0.250± | 0.153 | 0.719± | 0.414 |
| 10000 ppm | 35 | 48.0± 5.0 | 0.012± | 0.003 | 0.224± | 0.044 | 0.212± | 0.017 | 0.222± | 0.064 | 0.644± | 0.043 |
| 20000 ppm | 30 | 43.5± 8.3 | 0.010± | 0.004 | 0.202± | 0.035 | 0.207± | 0.028 | 0.261± | 0.148 | 0.719± | 0.491 |
| 40000 ppm | 25 | 42.9± 9.3 | 0.012± | 0.004 | 0.208± | 0.037 | 0.195± | 0.025** | 0.211± | 0.025 | 0.633± | 0.079 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of Animals | SPLEEN | | LIVER | | BRAIN | |
|------------|-------------------|--------|-------|--------|-------|--------|-------|
| Control | 34 | 0.214± | 0.528 | 1.883± | 0.615 | 0.458± | 0.015 |
| 10000 ppm | 35 | 0.119± | 0.197 | 1.910± | 0.686 | 0.453± | 0.016 |
| 20000 ppm | 30 | 0.189± | 0.263 | 2.060± | 0.907 | 0.455± | 0.015 |
| 40000 ppm | 25 | 0.123± | 0.110 | 1.741± | 0.466 | 0.449± | 0.019 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$ Test of Dunnett

(HCL040)

BAIS3

APPENDIX J 4

ORGAN WEIGHT, ABSOLUTE (TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105%)

PAGE : 3

| Group Name | NO. of Animals | Body Weight | ADRENALS | OVARIES | HEART | LUNGS | KIDNEYS |
|------------|----------------|-------------|--------------|--------------|--------------|---------------|----------------|
| Control | 29 | 32.7± 4.2 | 0.012± 0.003 | 0.035± 0.038 | 0.162± 0.019 | 0.196± 0.022 | 0.426± 0.044 |
| 10000 ppm | 24 | 31.5± 4.1 | 0.013± 0.003 | 0.093± 0.242 | 0.163± 0.020 | 0.212± 0.031 | 0.438± 0.037 |
| 20000 ppm | 19 | 32.5± 2.7 | 0.013± 0.003 | 0.058± 0.136 | 0.183± 0.075 | 0.223± 0.069* | 0.475± 0.090 |
| 40000 ppm | 28 | 31.4± 4.2 | 0.013± 0.004 | 0.076± 0.162 | 0.163± 0.022 | 0.221± 0.067* | 0.499± 0.181** |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS 3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE
UNIT: g

ORGAN WEIGHT:ABSOLUTE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | SPLEEN | | LIVER | | BRAIN | |
|------------|-------------------|--------|-------|--------|-------|--------|-------|
| Control | 29 | 0.151± | 0.167 | 1.464± | 0.232 | 0.487± | 0.070 |
| 10000 ppm | 24 | 0.174± | 0.147 | 1.453± | 0.166 | 0.478± | 0.018 |
| 20000 ppm | 19 | 0.175± | 0.103 | 1.752± | 0.619 | 0.481± | 0.024 |
| 40000 ppm | 28 | 0.211± | 0.163 | 1.767± | 1.110 | 0.470± | 0.024 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL040)

BAIS3

APPENDIX K 1

ORGAN WEIGHT, RELATIVE (TOW-YEAR STUDY: SUMMARY)

RAT : MALE

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of Animals | Body Weight (g) | ADRENALS | TESTES | HEART | LUNGS | KIDNEYS |
|------------|-------------------|--------------------|--------------|----------------|--------------|---------------|--------------|
| Control | 38 | 420± 50 | 0.023± 0.019 | 0.989± 0.337 | 0.298± 0.042 | 0.352± 0.047 | 0.756± 0.200 |
| 7500 ppm | 39 | 418± 34 | 0.020± 0.005 | 1.068± 0.352 | 0.296± 0.037 | 0.360± 0.058 | 0.758± 0.144 |
| 15000 ppm | 34 | 403± 38 | 0.043± 0.133 | 1.200± 0.362* | 0.298± 0.032 | 0.376± 0.045* | 0.771± 0.151 |
| 30000 ppm | 30 | 378± 40** | 0.022± 0.009 | 1.296± 0.425** | 0.305± 0.039 | 0.409± 0.154* | 0.744± 0.078 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS 3

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of Animals | SPLEEN | LIVER | BRAIN |
|------------|-------------------|--------------|--------------|----------------|
| Control | 38 | 0.282± 0.093 | 3.203± 0.543 | 0.490± 0.066 |
| 7500 ppm | 39 | 0.391± 0.553 | 3.287± 0.638 | 0.485± 0.041 |
| 15000 ppm | 34 | 0.304± 0.069 | 3.193± 0.394 | 0.503± 0.048 |
| 30000 ppm | 30 | 0.377± 0.312 | 3.277± 1.421 | 0.536± 0.059** |

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS 3

APPENDIX K 2

ORGAN WEIGHT, RELATIVE (TOW-YEAR STUDY: SUMMARY)

RAT : FEMALE

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | Body Weight (g) | ADRENALS | OVARIES | HEART | LUNGS | KIDNEYS |
|------------|-------------------|--------------------|----------------|--------------|----------------|----------------|----------------|
| Control | 47 | 294± 34 | 0.024± 0.003 | 0.058± 0.067 | 0.306± 0.033 | 0.353± 0.041 | 0.666± 0.068 |
| 7500 ppm | 39 | 293± 31 | 0.033± 0.038 | 0.072± 0.131 | 0.312± 0.035 | 0.395± 0.167 | 0.686± 0.123 |
| 15000 ppm | 35 | 276± 38* | 0.028± 0.008** | 0.158± 0.584 | 0.321± 0.050 | 0.385± 0.084 | 0.741± 0.122** |
| 30000 ppm | 14 | 258± 20** | 0.028± 0.003** | 0.052± 0.009 | 0.336± 0.030** | 0.421± 0.126** | 0.794± 0.061** |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS3

STUDY NO. : 0224
ANIMAL : RAT F344
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | SPLEEN | LIVER | BRAIN |
|------------|-------------------|--------------|--------------|----------------|
| Control | 47 | 0.220± 0.112 | 2.622± 0.363 | 0.635± 0.090 |
| 7500 ppm | 39 | 0.288± 0.370 | 2.611± 0.477 | 0.635± 0.084 |
| 15000 ppm | 35 | 0.246± 0.166 | 2.690± 0.485 | 0.676± 0.107 |
| 30000 ppm | 14 | 0.313± 0.358 | 2.784± 0.346 | 0.714± 0.051** |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS3

APPENDIX K 3

ORGAN WEIGHT, RELATIVE (TOW-YEAR STUDY: SUMMARY)

MOSUE : MALE

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 1

| Group Name | NO. of Animals | Body Weight (g) | ADRENALS | TESTES | HEART | LUNGS | KIDNEYS |
|------------|-------------------|--------------------|--------------|--------------|--------------|--------------|--------------|
| Control | 34 | 45.2± 8.5 | 0.031± 0.033 | 0.506± 0.153 | 0.518± 0.176 | 0.613± 0.601 | 1.699± 1.381 |
| 10000 ppm | 35 | 48.0± 5.0 | 0.024± 0.007 | 0.470± 0.103 | 0.447± 0.060 | 0.472± 0.174 | 1.354± 0.162 |
| 20000 ppm | 30 | 43.5± 8.3 | 0.024± 0.009 | 0.482± 0.140 | 0.491± 0.100 | 0.651± 0.542 | 1.708± 1.248 |
| 40000 ppm | 25 | 42.9± 9.3 | 0.029± 0.015 | 0.501± 0.106 | 0.474± 0.112 | 0.515± 0.123 | 1.528± 0.299 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : MALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 2

| Group Name | NO. of Animals | SPLEEN | LIVER | BRAIN |
|------------|-------------------|--------------|--------------|--------------|
| Control | 34 | 0.540± 1.476 | 4.370± 1.963 | 1.054± 0.233 |
| 10000 ppm | 35 | 0.273± 0.539 | 4.053± 1.695 | 0.954± 0.107 |
| 20000 ppm | 30 | 0.461± 0.641 | 5.087± 3.264 | 1.094± 0.280 |
| 40000 ppm | 25 | 0.307± 0.293 | 4.346± 1.957 | 1.104± 0.287 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS3

APPENDIX K 4

ORGAN WEIGHT, RELATIVE (TOW-YEAR STUDY: SUMMARY)

MOSUE : FEMALE

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 3

| Group Name | NO. of Animals | Body Weight (g) | ADRENALS | OVARIES | HEART | LUNGS | KIDNEYS |
|------------|----------------|-----------------|--------------|---------------|--------------|---------------|----------------|
| Control | 29 | 32.7± 4.2 | 0.038± 0.010 | 0.109± 0.123 | 0.502± 0.086 | 0.607± 0.094 | 1.318± 0.194 |
| 10000 ppm | 24 | 31.5± 4.1 | 0.042± 0.010 | 0.302± 0.812* | 0.527± 0.095 | 0.689± 0.152 | 1.415± 0.220 |
| 20000 ppm | 19 | 32.5± 2.7 | 0.040± 0.009 | 0.169± 0.368 | 0.565± 0.228 | 0.688± 0.210 | 1.469± 0.272* |
| 40000 ppm | 28 | 31.4± 4.2 | 0.042± 0.012 | 0.248± 0.557 | 0.523± 0.061 | 0.721± 0.263* | 1.642± 0.855** |

Significant difference : * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS3

STUDY NO. : 0225
ANIMAL : MOUSE BDF1
REPORT TYPE : A1
SEX : FEMALE
UNIT: %

ORGAN WEIGHT:RELATIVE (SUMMARY)
SURVIVAL ANIMALS (105W)

PAGE : 4

| Group Name | NO. of Animals | SPLEEN | LIVER | BRAIN |
|------------|-------------------|--------------|----------------|--------------|
| Control | 29 | 0.457± 0.480 | 4.528± 0.839 | 1.535± 0.492 |
| 10000 ppm | 24 | 0.572± 0.526 | 4.676± 0.743 | 1.544± 0.213 |
| 20000 ppm | 19 | 0.544± 0.317 | 5.397± 1.829 | 1.492± 0.146 |
| 40000 ppm | 28 | 0.694± 0.586 | 5.529± 2.579** | 1.514± 0.167 |

Significant difference ; * : $P \leq 0.05$ ** : $P \leq 0.01$

Test of Dunnett

(HCL042)

BAIS3