# Female BMD Results for Table: Liver Weight Absolute

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 10 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| Mean ± SD | 7.661 ± 0.788 | 7.626 ± 1.399 | 7.904 ± 0.801 | 8.436 ± 0.859 | 7.99 ± 0.988 | 8.054 ± 0.7 | 8.92 ± 1.069 | 9.278 ± 0.619 | 8.73 ± 0.833 | 9.378 ± 0.262 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | 0.031 | 51.333 | 647.314 | 425.62 | Exponential-M4 recommended as best-fitting model on the basis of the lowest AIC. |
| Hill | 0.555 | 42.327 | 23.753 | 3.734 |
| Exponential M2 (equivalent models include Exponential M3) | 0.029 | 51.53 | 675.627 | 458.479 |
| Exponential M4b (equivalent models include Exponential M5) | 0.598 | 41.973 | 24.928 | 7.768 |

a Constant variance case presented (BMDS Test 2 p-value = 0.174, BMDS Test 3 p-value = 0.174).

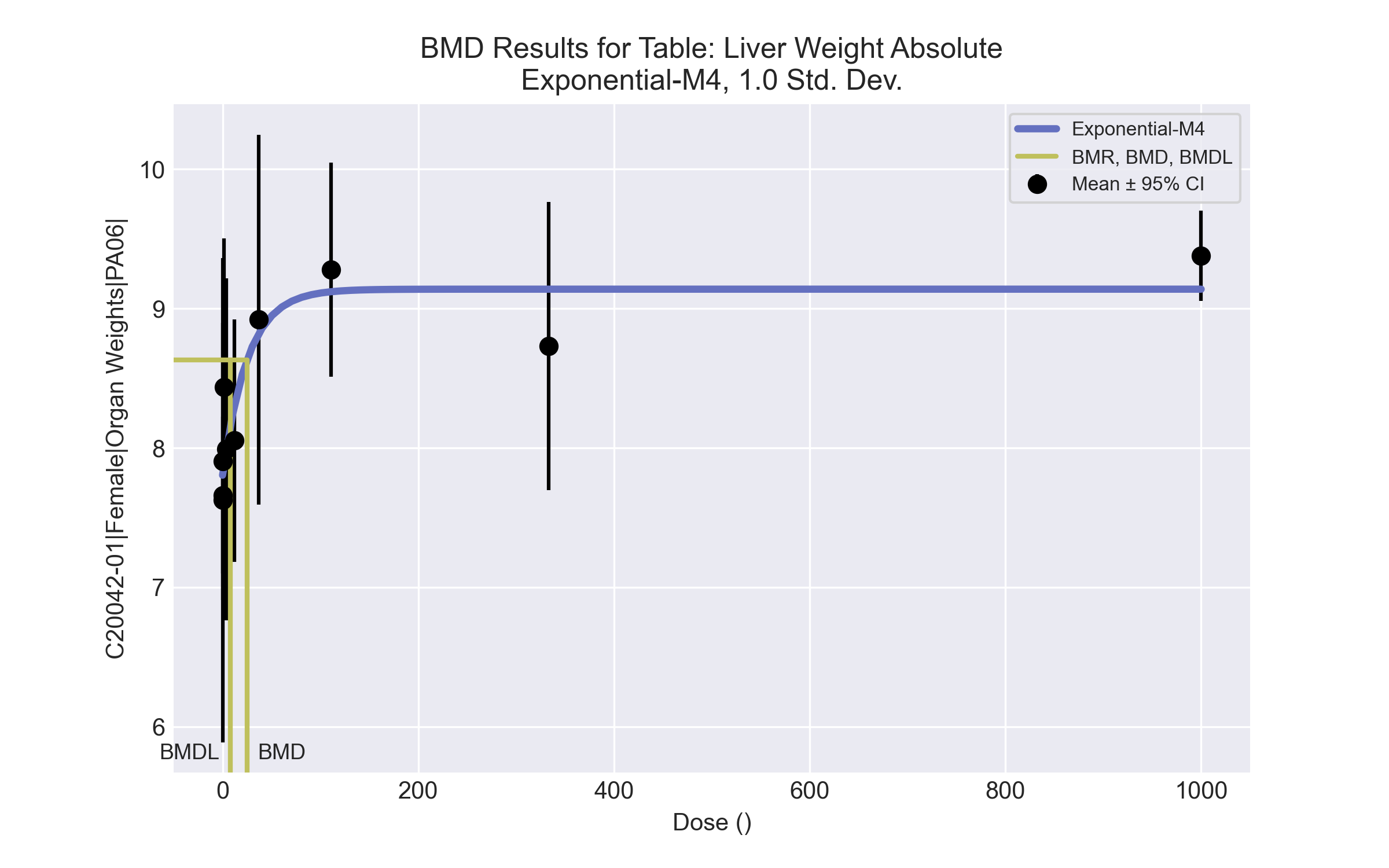
b Recommended model

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0314 < 0.1) |
| Hill | Valid | **Cautions**  • BMD/BMDL ratio is greater than threshold (6.36 > 5.0) |
| Exponential M2 (equivalent models include Exponential M3) | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0294 < 0.1) |
| Exponential M4a (equivalent models include Exponential M5) | Valid | - |

a Recommended model

## Recommended model



====================================================================   
 Exponential Model. (Version: 1.11; Date: 03/14/2017)   
 Input Data File: C:\Windows\TEMP\bmds-sa1w\_6wk.(d)   
 Gnuplot Plotting File:   
 Tue Apr 13 12:25:00 2021  
 ====================================================================   
  
 BMDS\_Model\_Run   
~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~  
   
 The form of the response function by Model:   
 Model 2: Y[dose] = a \* exp{sign \* b \* dose}  
 Model 3: Y[dose] = a \* exp{sign \* (b \* dose)^d}  
 Model 4: Y[dose] = a \* [c-(c-1) \* exp{-b \* dose}]  
 Model 5: Y[dose] = a \* [c-(c-1) \* exp{-(b \* dose)^d}]  
  
 Note: Y[dose] is the median response for exposure = dose;  
 sign = +1 for increasing trend in data;  
 sign = -1 for decreasing trend.  
  
 Model 2 is nested within Models 3 and 4.  
 Model 3 is nested within Model 5.  
 Model 4 is nested within Model 5.  
  
  
 Dependent variable = Response  
 Independent variable = Dose  
 Data are assumed to be distributed: normally  
 Variance Model: exp(lnalpha +rho \*ln(Y[dose]))  
 rho is set to 0.  
 A constant variance model is fit.  
  
 Total number of dose groups = 10  
 Total number of records with missing values = 0  
 Maximum number of iterations = 500  
 Relative Function Convergence has been set to: 1e-008  
 Parameter Convergence has been set to: 1e-008  
  
 MLE solution provided: Exact  
  
  
 Initial Parameter Values  
  
 Variable Model 4  
 -------- --------  
 lnalpha -0.482559   
 rho 0 Specified  
 a 7.2447   
 b 0.00196417   
 c 1.35919   
 d 1 Specified  
  
  
  
 Parameter Estimates  
  
 Variable Model 4 Std. Err.  
 -------- ------- ---------  
 lnalpha -0.382307 0.130107  
 a 7.80419 0.160982  
 b 0.0387084 0.0245564  
 c 1.17099 0.0360047  
  
 NC = No Convergence  
  
  
 Table of Stats From Input Data  
  
 Dose N Obs Mean Obs Std Dev  
 ----- --- ---------- -------------  
 0 10 7.661 0.788  
 0.15 5 7.626 1.399  
 0.5 5 7.904 0.801  
 1.4 5 8.436 0.8588  
 4 5 7.99 0.9876  
 12 5 8.054 0.7004  
 37 5 8.92 1.069  
 111 5 9.278 0.6188  
 333 5 8.73 0.8326  
 1000 5 9.378 0.2619  
  
  
 Estimated Values of Interest  
  
 Dose Est Mean Est Std Scaled Residual  
 ------ ---------- --------- ----------------  
 0 7.804 0.826 -0.5482  
 0.15 7.812 0.826 -0.5033  
 0.5 7.83 0.826 0.2009  
 1.4 7.875 0.826 1.52  
 4 7.996 0.826 -0.01518  
 12 8.3 0.826 -0.666  
 37 8.82 0.826 0.2707  
 111 9.12 0.826 0.4265  
 333 9.139 0.826 -1.106  
 1000 9.139 0.826 0.648  
  
  
  
 Other models for which likelihoods are calculated:  
  
 Model A1: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
  
 Model A2: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma(i)^2  
  
 Model A3: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = exp(lalpha + log(mean(i)) \* rho)  
  
 Model R: Yij = Mu + e(i)  
 Var{e(ij)} = Sigma^2  
  
  
 Likelihoods of Interest  
  
 Model Log(likelihood) DF AIC  
 ------- ----------------- ---- ------------  
 A1 -14.22964 11 50.45928  
 A2 -7.853379 20 55.70676  
 A3 -14.22964 11 50.45928  
 R -27.67333 2 59.34667  
 4 -16.98655 4 41.9731  
  
  
 Additive constant for all log-likelihoods = -50.54. This constant added to the  
 above values gives the log-likelihood including the term that does not  
 depend on the model parameters.  
  
  
 Explanation of Tests  
  
 Test 1: Does response and/or variances differ among Dose levels? (A2 vs. R)  
 Test 2: Are Variances Homogeneous? (A2 vs. A1)  
 Test 3: Are variances adequately modeled? (A2 vs. A3)  
  
 Test 6a: Does Model 4 fit the data? (A3 vs 4)  
  
  
 Tests of Interest  
  
 Test -2\*log(Likelihood Ratio) D. F. p-value  
 -------- ------------------------ ------ --------------  
 Test 1 39.64 18 0.002336  
 Test 2 12.75 9 0.1741  
 Test 3 12.75 9 0.1741  
 Test 6a 5.514 7 0.5975  
  
  
 The p-value for Test 1 is less than .05. There appears to be a  
 difference between response and/or variances among the dose  
 levels, it seems appropriate to model the data.  
  
 The p-value for Test 2 is greater than .1. A homogeneous  
 variance model appears to be appropriate here.  
  
 The p-value for Test 3 is greater than .1. The modeled  
 variance appears to be appropriate here.  
  
 The p-value for Test 6a is greater than .1. Model 4 seems  
 to adequately describe the data.  
  
  
 Benchmark Dose Computations:  
  
 Specified Effect = 1.000000  
  
 Risk Type = Estimated standard deviations from control  
  
 Confidence Level = 0.950000  
  
 BMD = 24.9281  
  
 BMDL = 7.76811  
  
 BMDU = 97.2949

# Female BMD Results for Table: Liver Weight Relative

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 10 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| Mean ± SD | 35.552 ± 1.626 | 35.557 ± 3.856 | 36.53 ± 1.442 | 36.913 ± 2.517 | 36.883 ± 1.533 | 36.799 ± 1.117 | 41.391 ± 4.457 | 42.008 ± 1.525 | 41.528 ± 2.61 | 44.404 ± 0.796 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | 3.3E-04 | 174.991 | 481.448 | 363.17 | No model was recommended as a best-fitting model. Doses were dropped until there were only 3 remaining dose-groups. |
| Hill | 0.41 | 156.187 | 18.65 | 6.452 |
| Exponential M2 (equivalent models include Exponential M3) | 2.8E-04 | 175.404 | 507.849 | 393.487 |
| Exponential M4 | 0.359 | 155.775 | 11.719 | 6.04 |
| Exponential M5 | 0.411 | 156.175 | 20.206 | 7.653 |

a Modeled variance case presented (BMDS Test 2 p-value = 0.002, BMDS Test 3 p-value = 9.7E-04).

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.0009702)  • Goodness of fit p-value is less than threshold (0.000327 < 0.1)  • Residual at lowest dose is greater than threshold (2.08 > 2.0)  • Ratio of modeled to actual stdev. at control is greater than threshold (2.02 > 1.5) |
| Hill | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.0009702) |
| Exponential M2 (equivalent models include Exponential M3) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.0009702)  • Goodness of fit p-value is less than threshold (0.000277 < 0.1)  • Residual at lowest dose is greater than threshold (2.1 > 2.0) |
| Exponential M4 | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.0009702) |
| Exponential M5 | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.0009702) |

## Recommended model

*No model was recommended as a best-fitting model.*

# Female BMD Results for Table: A/G Ratio

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 10 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| Mean ± SD | 2.385 ± 0.239 | 2.759 ± 0.098 | 2.364 ± 0.123 | 2.437 ± 0.087 | 2.499 ± 0.202 | 2.546 ± 0.103 | 2.26 ± 0.08 | 2.367 ± 0.139 | 2.455 ± 0.177 | 2.349 ± 0.159 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear | 1.6E-04 | -119.851 | -9999 | 2505.45 | No model was recommended as a best-fitting model. Doses were dropped until there were only 3 remaining dose-groups. |
| Polynomial 2° | 1.6E-04 | -119.851 | -9999 | 1565.98 |
| Polynomial 3° | 1.6E-04 | -119.851 | -9999 | 1315.41 |
| Polynomial 4° | 1.6E-04 | -119.851 | -9999 | 1515.29 |
| Polynomial 5° | 1.6E-04 | -119.851 | -9999 | -999 |
| Polynomial 6° | 1.6E-04 | -119.851 | -9999 | -999 |
| Polynomial 7° | 1.6E-04 | -119.851 | -9999 | -999 |
| Polynomial 8° | 1.6E-04 | -119.851 | -9999 | -999 |
| Power | 4.4E-04 | -122.393 | 1964 | 873.884 |
| Hill | 4.3E-04 | -121.921 | -999 | -999 |
| Exponential M2 | <0.0001 | -117.851 | 1 | -999 |
| Exponential M3 | <0.0001 | -99.743 | -1281.28 | -999 |
| Exponential M4 | <0.0001 | -115.851 | -999 | 0 |
| Exponential M5 | <0.0001 | -115.851 | -999 | 0 |

a Modeled variance case presented (BMDS Test 2 p-value = 0.059, BMDS Test 3 p-value = 0.073).

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear | Warning | **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): THE MODEL HAS PROBABLY NOT CONVERGED!!!; THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose)  • BMDL/high dose ratio is greater than threshold (2.51 > 1.0) |
| Polynomial 2° | Warning | **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose)  • BMDL/high dose ratio is greater than threshold (1.57 > 1.0) |
| Polynomial 3° | Warning | **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): BMD = 100\*(maximum dose)  • BMDL/high dose ratio is greater than threshold (1.32 > 1.0) |
| Polynomial 4° | Warning | **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): BMD = 100\*(maximum dose)  • BMDL/high dose ratio is greater than threshold (1.52 > 1.0) |
| Polynomial 5° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): BMD = 100\*(maximum dose); BMDL computation failed. |
| Polynomial 6° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose); BMDL computation failed. |
| Polynomial 7° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose); BMDL computation failed. |
| Polynomial 8° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000162 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose); BMDL computation failed. |
| Power | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000443 < 0.1)  • BMD/high dose ratio is greater than threshold (1.96 > 1.0) |
| Hill | Failure | **Failures**  • BMD does not exist  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325)  • Goodness of fit p-value is less than threshold (0.000432 < 0.1)  **Cautions**  • Warning(s): BMR value is not in the range of the mean function |
| Exponential M2 | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.07325) |
| Exponential M3 | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325) |
| Exponential M4 | Failure | **Failures**  • BMD does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325) |
| Exponential M5 | Failure | **Failures**  • BMD does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.07325) |

## Recommended model

*No model was recommended as a best-fitting model.*

# Female BMD Results for Table: Alkaline phosphatase

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 10 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| Mean ± SD | 201.3 ± 41.358 | 198.8 ± 26.593 | 216.2 ± 86.803 | 210 ± 44.565 | 218.4 ± 22.568 | 208.4 ± 16.441 | 222.8 ± 55.979 | 240 ± 20.748 | 241.4 ± 12.759 | 272.2 ± 40.678 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 8°) | 0.283 | 466.971 | 644.753 | 427.604 | No model was recommended as a best-fitting model. Doses were dropped until there were only 3 remaining dose-groups. |
| Polynomial 7° | 0.283 | 466.971 | 644.752 | 427.604 |
| Hill | 0.526 | 465.333 | 222.224 | -999 |
| Exponential M2 (equivalent models include Exponential M3) | 0.261 | 467.278 | 681.136 | 472.905 |
| Exponential M4 (equivalent models include Exponential M5) | 0.465 | 465.885 | 225.225 | 47.683 |

a Modeled variance case presented (BMDS Test 2 p-value = 4.1E-04, BMDS Test 3 p-value = 0.003).

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 8°) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.003117) |
| Polynomial 7° | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.003117) |
| Hill | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.003117)  **Cautions**  • Warning(s): BMDL computation failed. |
| Exponential M2 (equivalent models include Exponential M3) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.003117) |
| Exponential M4 (equivalent models include Exponential M5) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.003117) |

## Recommended model

*No model was recommended as a best-fitting model.*

# Female BMD Results for Table: Cholesterol

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 10 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| Mean ± SD | 105 ± 16.607 | 97.8 ± 12.95 | 100.8 ± 15.802 | 88.6 ± 10.502 | 101 ± 8.515 | 109 ± 27.359 | 123.8 ± 23.264 | 105.6 ± 12.341 | 120.8 ± 13.498 | 162.4 ± 19.087 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linearb (equivalent models include Power) | 0.135 | 373.284 | 288.242 | 226.543 | Linear recommended as best-fitting model on the basis of the lowest AIC. |
| Polynomial 2° | 0.089 | 375.282 | 292.517 | 226.555 |
| Polynomial 3° | 0.089 | 375.274 | 299.479 | 226.625 |
| Polynomial 4° | 0.089 | 375.272 | 300.989 | 226.65 |
| Polynomial 5° | 0.089 | 375.271 | 301.38 | 226.658 |
| Polynomial 6° | 0.089 | 375.271 | 301.491 | 226.66 |
| Polynomial 7° | 0.089 | 375.271 | 301.524 | 226.661 |
| Polynomial 8° | 0.054 | 377.27 | 301.532 | 226.661 |
| Hill | 0.089 | 375.284 | 287.68 | 225.981 |
| Exponential M2 (equivalent models include Exponential M3) | 0.131 | 373.386 | 336.592 | 274.666 |
| Exponential M4 | 0.089 | 375.284 | 287.087 | 149.15 |
| Exponential M5 | 0.089 | 375.285 | 286.834 | 149.146 |

a Constant variance case presented (BMDS Test 2 p-value = 0.246, BMDS Test 3 p-value = 0.246).

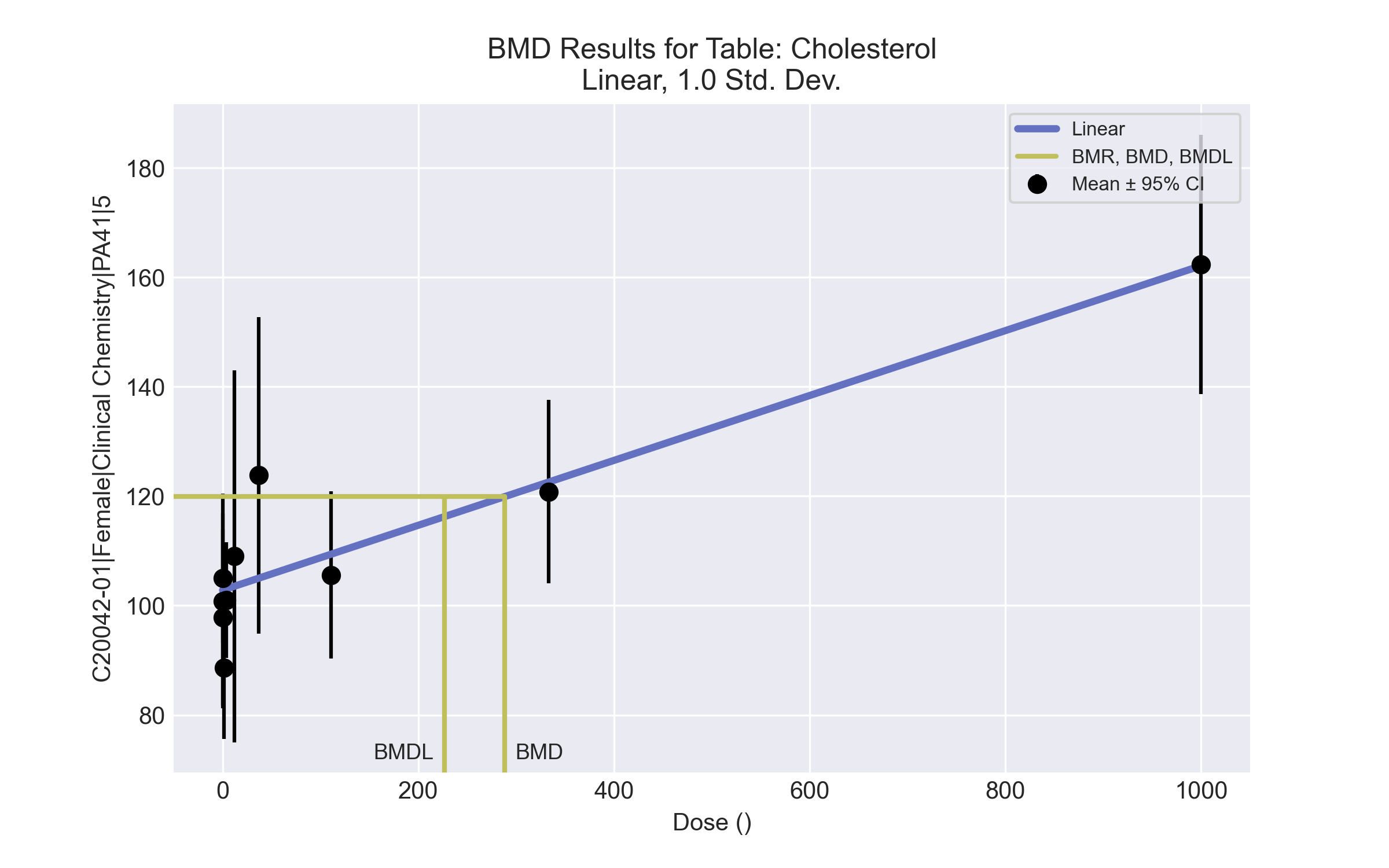
b Recommended model

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Lineara (equivalent models include Power) | Valid | - |
| Polynomial 2° | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0887 < 0.1) |
| Polynomial 3° | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0889 < 0.1) |
| Polynomial 4° | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.089 < 0.1) |
| Polynomial 5° | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.089 < 0.1) |
| Polynomial 6° | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.089 < 0.1) |
| Polynomial 7° | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.089 < 0.1) |
| Polynomial 8° | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0542 < 0.1) |
| Hill | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0886 < 0.1) |
| Exponential M2 (equivalent models include Exponential M3) | Valid | - |
| Exponential M4 | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0886 < 0.1) |
| Exponential M5 | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.0886 < 0.1) |

a Recommended model

## Recommended model



====================================================================   
 Polynomial Model. (Version: 2.21; Date: 03/14/2017)   
 Input Data File: C:\Windows\TEMP\bmds-m3\_o4mnn.(d)   
 Gnuplot Plotting File: C:\Windows\TEMP\bmds-m3\_o4mnn.plt  
 Tue Apr 13 13:46:42 2021  
 ====================================================================   
  
 BMDS\_Model\_Run   
~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~  
   
 The form of the response function is:   
  
 Y[dose] = beta\_0 + beta\_1\*dose + beta\_2\*dose^2 + ...  
  
  
 Dependent variable = Response  
 Independent variable = Dose  
 rho is set to 0  
 The polynomial coefficients are restricted to be positive  
 A constant variance model is fit  
  
 Total number of dose groups = 10  
 Total number of records with missing values = 0  
 Maximum number of iterations = 500  
 Relative Function Convergence has been set to: 1e-008  
 Parameter Convergence has been set to: 1e-008  
  
  
  
 Default Initial Parameter Values   
 alpha = 285.262  
 rho = 0 Specified  
 beta\_0 = 102.533  
 beta\_1 = 0  
  
  
 Asymptotic Correlation Matrix of Parameter Estimates  
  
 ( \*\*\* The model parameter(s) -rho   
 have been estimated at a boundary point, or have been specified by the user,  
 and do not appear in the correlation matrix )  
  
 alpha beta\_0 beta\_1  
  
 alpha 1 1.6e-006 4.7e-007  
  
 beta\_0 1.6e-006 1 -0.43  
  
 beta\_1 4.7e-007 -0.43 1  
  
  
  
 Parameter Estimates  
  
 95.0% Wald Confidence Interval  
 Variable Estimate Std. Err. Lower Conf. Limit Upper Conf. Limit  
 alpha 292.33 55.7452 183.072 401.589  
 beta\_0 102.807 2.54848 97.8124 107.802  
 beta\_1 0.0593171 0.00796986 0.0436965 0.0749377  
  
  
  
 Table of Data and Estimated Values of Interest  
  
 Dose N Obs Mean Est Mean Obs Std Dev Est Std Dev Scaled Res.  
------ --- -------- -------- ----------- ----------- ----------  
  
 0 10 105 103 16.6 17.1 0.406  
 0.15 5 97.8 103 12.9 17.1 -0.656  
 0.5 5 101 103 15.8 17.1 -0.266  
 1.4 5 88.6 103 10.5 17.1 -1.87  
 4 5 101 103 8.51 17.1 -0.267  
 12 5 109 104 27.4 17.1 0.717  
 37 5 124 105 23.3 17.1 2.46  
 111 5 106 109 12.3 17.1 -0.496  
 333 5 121 123 13.5 17.1 -0.23  
 1000 5 162 162 19.1 17.1 0.036  
  
  
  
 Model Descriptions for likelihoods calculated  
  
  
 Model A1: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
  
 Model A2: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma(i)^2  
  
 Model A3: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
 Model A3 uses any fixed variance parameters that  
 were specified by the user  
  
 Model R: Yi = Mu + e(i)  
 Var{e(i)} = Sigma^2  
  
  
 Likelihoods of Interest  
  
 Model Log(likelihood) # Param's AIC  
 A1 -177.450299 11 376.900598  
 A2 -171.725501 20 383.451001  
 A3 -177.450299 11 376.900598  
 fitted -183.641824 3 373.283649  
 R -202.801549 2 409.603099  
  
  
 Explanation of Tests   
  
 Test 1: Do responses and/or variances differ among Dose levels?   
 (A2 vs. R)  
 Test 2: Are Variances Homogeneous? (A1 vs A2)  
 Test 3: Are variances adequately modeled? (A2 vs. A3)  
 Test 4: Does the Model for the Mean Fit? (A3 vs. fitted)  
 (Note: When rho=0 the results of Test 3 and Test 2 will be the same.)  
  
 Tests of Interest   
  
 Test -2\*log(Likelihood Ratio) Test df p-value   
  
 Test 1 62.1521 18 <.0001  
 Test 2 11.4496 9 0.2461  
 Test 3 11.4496 9 0.2461  
 Test 4 12.3831 8 0.1349  
  
The p-value for Test 1 is less than .05. There appears to be a  
difference between response and/or variances among the dose levels  
It seems appropriate to model the data  
  
The p-value for Test 2 is greater than .1. A homogeneous variance   
model appears to be appropriate here  
  
  
The p-value for Test 3 is greater than .1. The modeled variance appears   
 to be appropriate here  
  
The p-value for Test 4 is greater than .1. The model chosen seems   
to adequately describe the data  
   
  
 Benchmark Dose Computation  
  
Specified effect = 1  
  
Risk Type = Estimated standard deviations from the control mean  
  
Confidence level = 0.95  
  
 BMD = 288.242  
  
  
 BMDL = 226.543  
  
  
 BMDU = 394.641

# Female BMD Results for Table: Globulin (measured)

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111a | 333a | 1000a |
| N | 10 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| Mean ± SD | 1.93 ± 0.226 | 1.66 ± 0.055 | 1.88 ± 0.11 | 1.88 ± 0.084 | 1.86 ± 0.182 | 1.84 ± 0.114 | 2.04 ± 0.207 | 1.94 ± 0.114 | 1.9 ± 0.2 | 1.98 ± 0.11 |

a Dose group removed in BMD modeling session

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear | 1.3E-04 | -90.762 | -9999 | 96.355 | Hill recommended as best-fitting model on the basis of the lowest AIC. |
| Polynomial 2° | 1.3E-04 | -90.762 | -9999 | 38.406 |
| Polynomial 3° | 1.3E-04 | -90.762 | -9999 | 37.074 |
| Polynomial 4° | 1.3E-04 | -90.762 | -9999 | 36.685 |
| Polynomial 5° | 1.3E-04 | -90.762 | -9999 | -999 |
| Polynomial 6° | 1.3E-04 | -90.762 | -9999 | -999 |
| Polynomial 7° | 1.3E-04 | -90.762 | -9999 | -999 |
| Polynomial 8° | <0.0001 | 396.005 | -9999 | -999 |
| Power | 1.8E-04 | -91.749 | 36.758 | 21.342 |
| Hillb | <0.0001 | -89.749 | 36.678 | 13.695 |
| Exponential M2 | <0.0001 | -88.762 | 1 | -999 |
| Exponential M3 | <0.0001 | -88.403 | 50.384 | 38.023 |
| Exponential M4 | <0.0001 | -86.762 | -999 | 0 |
| Exponential M5 | <0.0001 | -86.762 | -999 | 0 |

a Modeled variance case presented (BMDS Test 2 p-value = 0.012, BMDS Test 3 p-value = 0.565).

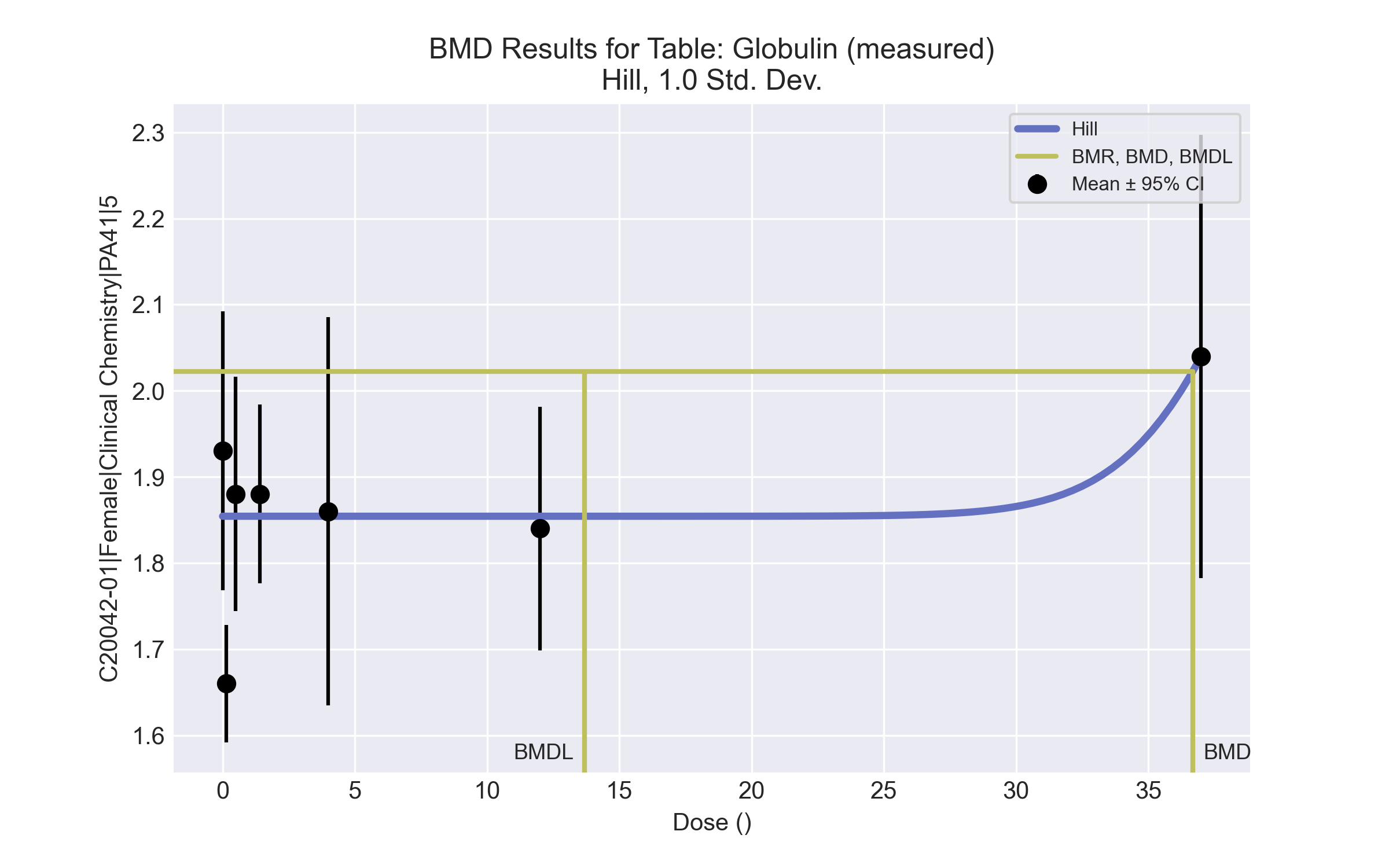
b Recommended model

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear | Warning | **Warnings**  • Residual of Interest does not exist  • Goodness of fit p-value is less than threshold (0.000128 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose)  • BMDL/high dose ratio is greater than threshold (2.6 > 1.0) |
| Polynomial 2° | Warning | **Warnings**  • Residual of Interest does not exist  • Goodness of fit p-value is less than threshold (0.000128 < 0.1)  **Cautions**  • Warning(s): BMD = 100\*(maximum dose)  • BMDL/high dose ratio is greater than threshold (1.04 > 1.0) |
| Polynomial 3° | Warning | **Warnings**  • Residual of Interest does not exist  • Goodness of fit p-value is less than threshold (0.000128 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose)  • BMDL/high dose ratio is greater than threshold (1.0 > 1.0) |
| Polynomial 4° | Warning | **Warnings**  • Residual of Interest does not exist  • Goodness of fit p-value is less than threshold (0.000128 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose) |
| Polynomial 5° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Goodness of fit p-value is less than threshold (0.000128 < 0.1)  **Cautions**  • Warning(s): BMD = 100\*(maximum dose); BMDL computation failed. |
| Polynomial 6° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Goodness of fit p-value is less than threshold (0.000128 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose); BMDL computation failed. |
| Polynomial 7° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Goodness of fit p-value is less than threshold (0.000128 < 0.1)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose); BMDL computation failed. |
| Polynomial 8° | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Ratio of modeled to actual stdev. at control is greater than threshold (4.51 > 1.5)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose); BMDL computation failed. |
| Power | Warning | **Warnings**  • Goodness of fit p-value is less than threshold (0.000175 < 0.1)  **Cautions**  • Warning(s): THE MODEL HAS PROBABLY NOT CONVERGED!!!; THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED! |
| Hilla | Valid | - |
| Exponential M2 | Failure | **Failures**  • BMDL does not exist |
| Exponential M3 | Warning | **Warnings**  • BMD/high dose ratio is greater than threshold (1.36 > 1.0)  **Cautions**  • BMDL/high dose ratio is greater than threshold (1.03 > 1.0) |
| Exponential M4 | Failure | **Failures**  • BMD does not exist  **Warnings**  • Residual of Interest does not exist |
| Exponential M5 | Failure | **Failures**  • BMD does not exist  **Warnings**  • Residual of Interest does not exist |

a Recommended model

## Recommended model



====================================================================   
 Hill Model. (Version: 2.18; Date: 03/14/2017)   
 Input Data File: C:\Windows\TEMP\bmds-9grtlgid.(d)   
 Gnuplot Plotting File: C:\Windows\TEMP\bmds-9grtlgid.plt  
 Tue Apr 13 13:47:02 2021  
 ====================================================================   
  
 BMDS\_Model\_Run   
~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~  
   
 The form of the response function is:   
  
 Y[dose] = intercept + v\*dose^n/(k^n + dose^n)  
  
  
 Dependent variable = Response  
 Independent variable = Dose  
 Power parameter restricted to be greater than 1  
 The variance is to be modeled as Var(i) = exp(lalpha + rho \* ln(mean(i)))  
  
 Total number of dose groups = 7  
 Total number of records with missing values = 0  
 Maximum number of iterations = 500  
 Relative Function Convergence has been set to: 1e-008  
 Parameter Convergence has been set to: 1e-008  
  
  
  
 Default Initial Parameter Values   
 lalpha = -3.59633  
 rho = 0  
 intercept = 1.93  
 v = 0.11  
 n = 1  
 k = 43.875  
  
  
 Asymptotic Correlation Matrix of Parameter Estimates  
  
 lalpha rho intercept v n k  
  
 lalpha 1 -1 -0.053 0.00025 -0.00051 0.0003  
  
 rho -1 1 0.047 -0.00024 0.00051 -0.0003  
  
 intercept -0.053 0.047 1 -0.00017 0.00029 -3.6e-005  
  
 v 0.00025 -0.00024 -0.00017 1 -0.48 0.94  
  
 n -0.00051 0.00051 0.00029 -0.48 1 -0.74  
  
 k 0.0003 -0.0003 -3.6e-005 0.94 -0.74 1  
  
  
  
 Parameter Estimates  
  
 95.0% Wald Confidence Interval  
 Variable Estimate Std. Err. Lower Conf. Limit Upper Conf. Limit  
 lalpha -4.85517 4.50401 -13.6829 3.97253  
 rho 2.08355 7.14672 -11.9238 16.0909  
 intercept 1.85429 0.028383 1.79866 1.90992  
 v 0.878941 517.356 -1013.12 1014.88  
 n 14.4172 3986.46 -7798.9 7827.73  
 k 40.5395 2738.75 -5327.31 5408.39  
  
  
  
 Table of Data and Estimated Values of Interest  
  
 Dose N Obs Mean Est Mean Obs Std Dev Est Std Dev Scaled Res.  
------ --- -------- -------- ----------- ----------- ----------  
  
 0 10 1.93 1.85 0.226 0.168 1.43  
 0.15 5 1.66 1.85 0.0548 0.168 -2.59  
 0.5 5 1.88 1.85 0.11 0.168 0.342  
 1.4 5 1.88 1.85 0.0837 0.168 0.342  
 4 5 1.86 1.85 0.182 0.168 0.0761  
 12 5 1.84 1.85 0.114 0.168 -0.19  
 37 5 2.04 2.04 0.207 0.185 1.39e-006  
  
  
  
 Model Descriptions for likelihoods calculated  
  
  
 Model A1: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
  
 Model A2: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma(i)^2  
  
 Model A3: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = exp(lalpha + rho\*ln(Mu(i)))  
 Model A3 uses any fixed variance parameters that  
 were specified by the user  
  
 Model R: Yi = Mu + e(i)  
 Var{e(i)} = Sigma^2  
  
  
 Likelihoods of Interest  
  
 Model Log(likelihood) # Param's AIC  
 A1 55.774020 8 -95.548039  
 A2 63.967982 14 -99.935965  
 A3 62.021623 9 -106.043247  
 fitted 50.874380 6 -89.748759  
 R 48.380880 2 -92.761760  
  
  
 Explanation of Tests   
  
 Test 1: Do responses and/or variances differ among Dose levels?   
 (A2 vs. R)  
 Test 2: Are Variances Homogeneous? (A1 vs A2)  
 Test 3: Are variances adequately modeled? (A2 vs. A3)  
 Test 4: Does the Model for the Mean Fit? (A3 vs. fitted)  
 (Note: When rho=0 the results of Test 3 and Test 2 will be the same.)  
  
 Tests of Interest   
  
 Test -2\*log(Likelihood Ratio) Test df p-value   
  
 Test 1 31.1742 12 0.001853  
 Test 2 16.3879 6 0.01182  
 Test 3 3.89272 5 0.565  
 Test 4 22.2945 3 <.0001  
  
The p-value for Test 1 is less than .05. There appears to be a  
difference between response and/or variances among the dose levels  
It seems appropriate to model the data  
  
The p-value for Test 2 is less than .1. A non-homogeneous variance   
model appears to be appropriate  
  
The p-value for Test 3 is greater than .1. The modeled variance appears   
 to be appropriate here  
  
The p-value for Test 4 is less than .1. You may want to try a different   
model  
   
  
 Benchmark Dose Computation  
  
Specified effect = 1  
  
Risk Type = Estimated standard deviations from the control mean   
  
Confidence level = 0.95  
  
 BMD = 36.6778  
  
 BMDL = 13.6954  
  
 BMDU = 1.369e+007

# Female BMD Results for Table: Eosinophil count

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 8 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 4 | 5 |
| Mean ± SD | 0.083 ± 0.029 | 0.078 ± 0.046 | 0.08 ± 0.031 | 0.092 ± 0.039 | 0.068 ± 0.042 | 0.05 ± 0.016 | 0.042 ± 0.027 | 0.076 ± 0.047 | 0.03 ± 0.022 | 0.044 ± 0.028 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | 0.105 | -291.4 | 1000.14 | 558.717 | Exponential-M5 recommended as best-fitting model on the basis of the lowest BMDL. |
| Hill | 0.414 | -294.524 | 5.053 | -999 |
| Exponential M2 (equivalent models include Exponential M3) | 0.126 | -292 | 820.54 | 281.561 |
| Exponential M4 | 0.45 | -295.805 | 17.001 | 3.067 |
| Exponential M5b | 0.414 | -294.524 | 4.428 | 1.564 |

a Constant variance case presented (BMDS Test 2 p-value = 0.397, BMDS Test 3 p-value = 0.397).

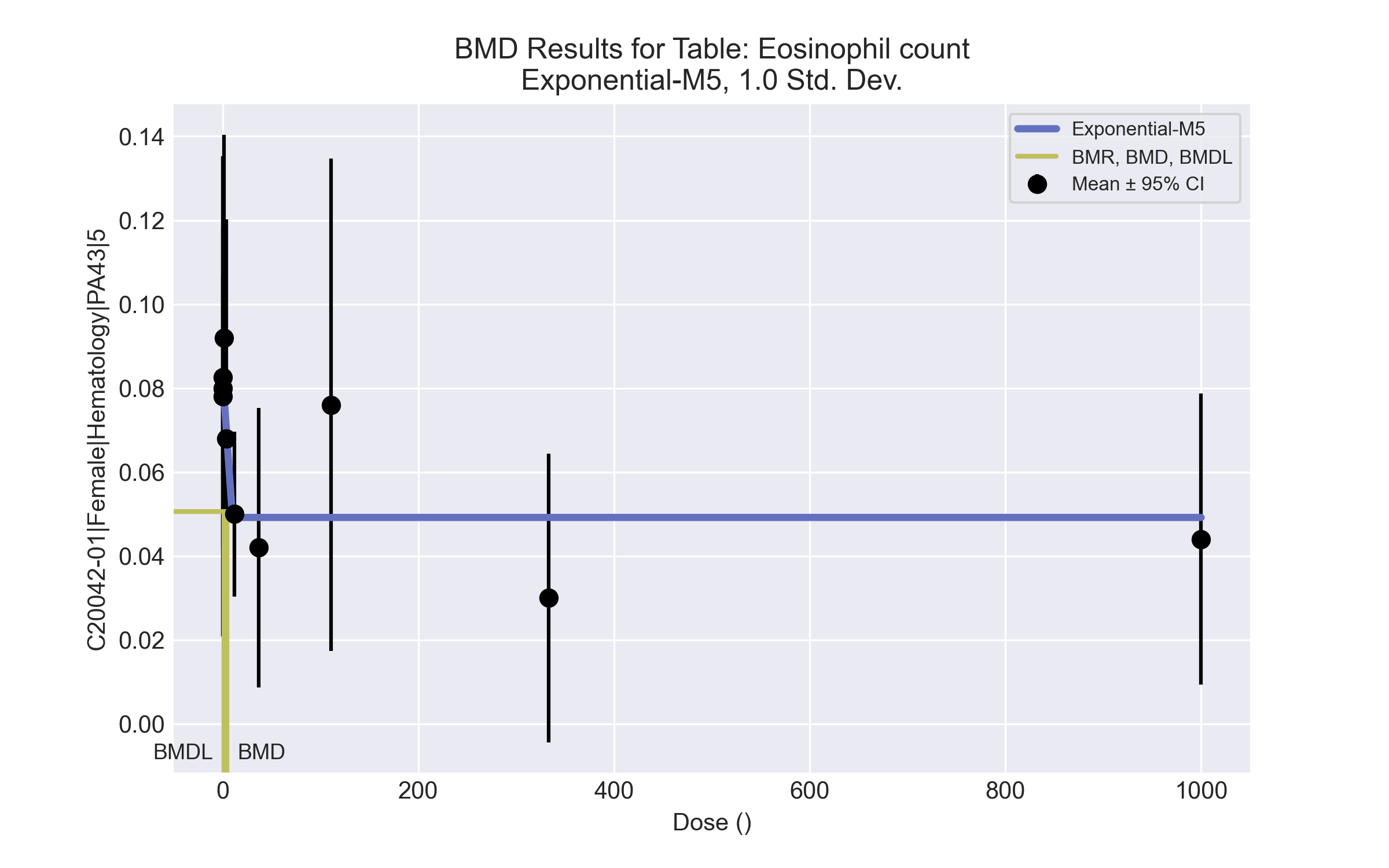
b Recommended model

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | Warning | **Warnings**  • BMD/high dose ratio is greater than threshold (1.0 > 1.0) |
| Hill | Failure | **Failures**  • BMDL does not exist  **Cautions**  • Warning(s): BMDL computation failed. |
| Exponential M2 (equivalent models include Exponential M3) | Valid | - |
| Exponential M4 | Valid | **Cautions**  • BMD/BMDL ratio is greater than threshold (5.54 > 5.0) |
| Exponential M5a | Valid | - |

a Recommended model

## Recommended model



====================================================================   
 Exponential Model. (Version: 1.11; Date: 03/14/2017)   
 Input Data File: C:\Windows\TEMP\bmds-n3ygn080.(d)   
 Gnuplot Plotting File:   
 Tue Apr 13 13:58:21 2021  
 ====================================================================   
  
 BMDS\_Model\_Run   
~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~  
   
 The form of the response function by Model:   
 Model 2: Y[dose] = a \* exp{sign \* b \* dose}  
 Model 3: Y[dose] = a \* exp{sign \* (b \* dose)^d}  
 Model 4: Y[dose] = a \* [c-(c-1) \* exp{-b \* dose}]  
 Model 5: Y[dose] = a \* [c-(c-1) \* exp{-(b \* dose)^d}]  
  
 Note: Y[dose] is the median response for exposure = dose;  
 sign = +1 for increasing trend in data;  
 sign = -1 for decreasing trend.  
  
 Model 2 is nested within Models 3 and 4.  
 Model 3 is nested within Model 5.  
 Model 4 is nested within Model 5.  
  
  
 Dependent variable = Response  
 Independent variable = Dose  
 Data are assumed to be distributed: normally  
 Variance Model: exp(lnalpha +rho \*ln(Y[dose]))  
 rho is set to 0.  
 A constant variance model is fit.  
  
 Total number of dose groups = 10  
 Total number of records with missing values = 0  
 Maximum number of iterations = 500  
 Relative Function Convergence has been set to: 1e-008  
 Parameter Convergence has been set to: 1e-008  
  
 MLE solution provided: Exact  
  
  
 Initial Parameter Values  
  
 Variable Model 5  
 -------- --------  
 lnalpha -6.97312   
 rho 0 Specified  
 a 0.0966   
 b 0.00256643   
 c 0.29577   
 d 1   
  
  
  
 Parameter Estimates  
  
 Variable Model 5 Std. Err.  
 -------- ------- ---------  
 lnalpha -6.85622 0.000206487  
 a 0.0830434 0.0067659  
 b 0.242104 2.3972  
 c 0.592059 0.0932113  
 d 16.5946 5119.88  
  
 NC = No Convergence  
  
  
 Table of Stats From Input Data  
  
 Dose N Obs Mean Obs Std Dev  
 ----- --- ---------- -------------  
 0 8 0.0825 0.02916  
 0.15 5 0.078 0.04604  
 0.5 5 0.08 0.03082  
 1.4 5 0.092 0.03899  
 4 5 0.068 0.04207  
 12 5 0.05 0.01581  
 37 5 0.042 0.02683  
 111 5 0.076 0.04722  
 333 4 0.03 0.0216  
 1000 5 0.044 0.02793  
  
  
 Estimated Values of Interest  
  
 Dose Est Mean Est Std Scaled Residual  
 ------ ---------- --------- ----------------  
 0 0.08304 0.03245 -0.04737  
 0.15 0.08304 0.03245 -0.3476  
 0.5 0.08304 0.03245 -0.2097  
 1.4 0.08304 0.03245 0.6172  
 4 0.068 0.03245 4.391e-006  
 12 0.04917 0.03245 0.05743  
 37 0.04917 0.03245 -0.4939  
 111 0.04917 0.03245 1.849  
 333 0.04917 0.03245 -1.181  
 1000 0.04917 0.03245 -0.356  
  
  
  
 Other models for which likelihoods are calculated:  
  
 Model A1: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
  
 Model A2: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma(i)^2  
  
 Model A3: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = exp(lalpha + log(mean(i)) \* rho)  
  
 Model R: Yij = Mu + e(i)  
 Var{e(ij)} = Sigma^2  
  
  
 Likelihoods of Interest  
  
 Model Log(likelihood) DF AIC  
 ------- ----------------- ---- ------------  
 A1 155.3012 11 -288.6024  
 A2 160.0273 20 -280.0546  
 A3 155.3012 11 -288.6024  
 R 146.5308 2 -289.0616  
 5 152.2618 5 -294.5236  
  
  
 Additive constant for all log-likelihoods = -47.78. This constant added to the  
 above values gives the log-likelihood including the term that does not  
 depend on the model parameters.  
  
  
 Explanation of Tests  
  
 Test 1: Does response and/or variances differ among Dose levels? (A2 vs. R)  
 Test 2: Are Variances Homogeneous? (A2 vs. A1)  
 Test 3: Are variances adequately modeled? (A2 vs. A3)  
  
 Test 7a: Does Model 5 fit the data? (A3 vs 5)  
  
  
 Tests of Interest  
  
 Test -2\*log(Likelihood Ratio) D. F. p-value  
 -------- ------------------------ ------ --------------  
 Test 1 26.99 18 0.07913  
 Test 2 9.452 9 0.3966  
 Test 3 9.452 9 0.3966  
 Test 7a 6.079 6 0.4144  
  
  
 The p-value for Test 1 is greater than .05. There may not be a  
 diffence between responses and/or variances among the dose levels  
 Modelling the data with a dose/response curve may not be appropriate.  
  
 The p-value for Test 2 is greater than .1. A homogeneous  
 variance model appears to be appropriate here.  
  
 The p-value for Test 3 is greater than .1. The modeled  
 variance appears to be appropriate here.  
  
 The p-value for Test 7a is greater than .1. Model 5 seems  
 to adequately describe the data.  
  
  
 Benchmark Dose Computations:  
  
 Specified Effect = 1.000000  
  
 Risk Type = Estimated standard deviations from control  
  
 Confidence Level = 0.950000  
  
 BMD = 4.4275  
  
 BMDL = 1.5636  
  
 BMDU = 1e+007

# Female BMD Results for Table: Mean Cell Hemoglobin

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000a |
| N | 8 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 4 | 5 |
| Mean ± SD | 17.863 ± 0.358 | 18.18 ± 0.455 | 18.08 ± 0.37 | 17.76 ± 0.23 | 17.66 ± 0.27 | 17.88 ± 0.277 | 17.82 ± 0.356 | 17.78 ± 0.396 | 17.45 ± 0.311 | 17.74 ± 0.483 |

a Dose group removed in BMD modeling session

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 2°) | 0.256 | -48.185 | 252.318 | 151.291 | Exponential-M2 recommended as best-fitting model on the basis of the lowest AIC. |
| Polynomial 3° | 0.177 | -46.186 | 255.577 | 151.301 |
| Polynomial 4° | 0.177 | -46.186 | 258.03 | 151.311 |
| Polynomial 5° | 0.177 | -46.187 | 259.675 | 151.314 |
| Polynomial 6° | 0.177 | -46.187 | 260.855 | 151.316 |
| Polynomial 7° | 0.177 | -46.187 | 261.743 | 151.316 |
| Polynomial 8° | 0.177 | -46.187 | 262.433 | 151.316 |
| Hill | 0.177 | -46.185 | 251.156 | -999 |
| Exponential M2b (equivalent models include Exponential M3) | 0.257 | -48.185 | 251.686 | 149.752 |
| Exponential M4 (equivalent models include Exponential M5) | 0.177 | -46.185 | 251.268 | 1.333 |

a Constant variance case presented (BMDS Test 2 p-value = 0.887, BMDS Test 3 p-value = 0.887).

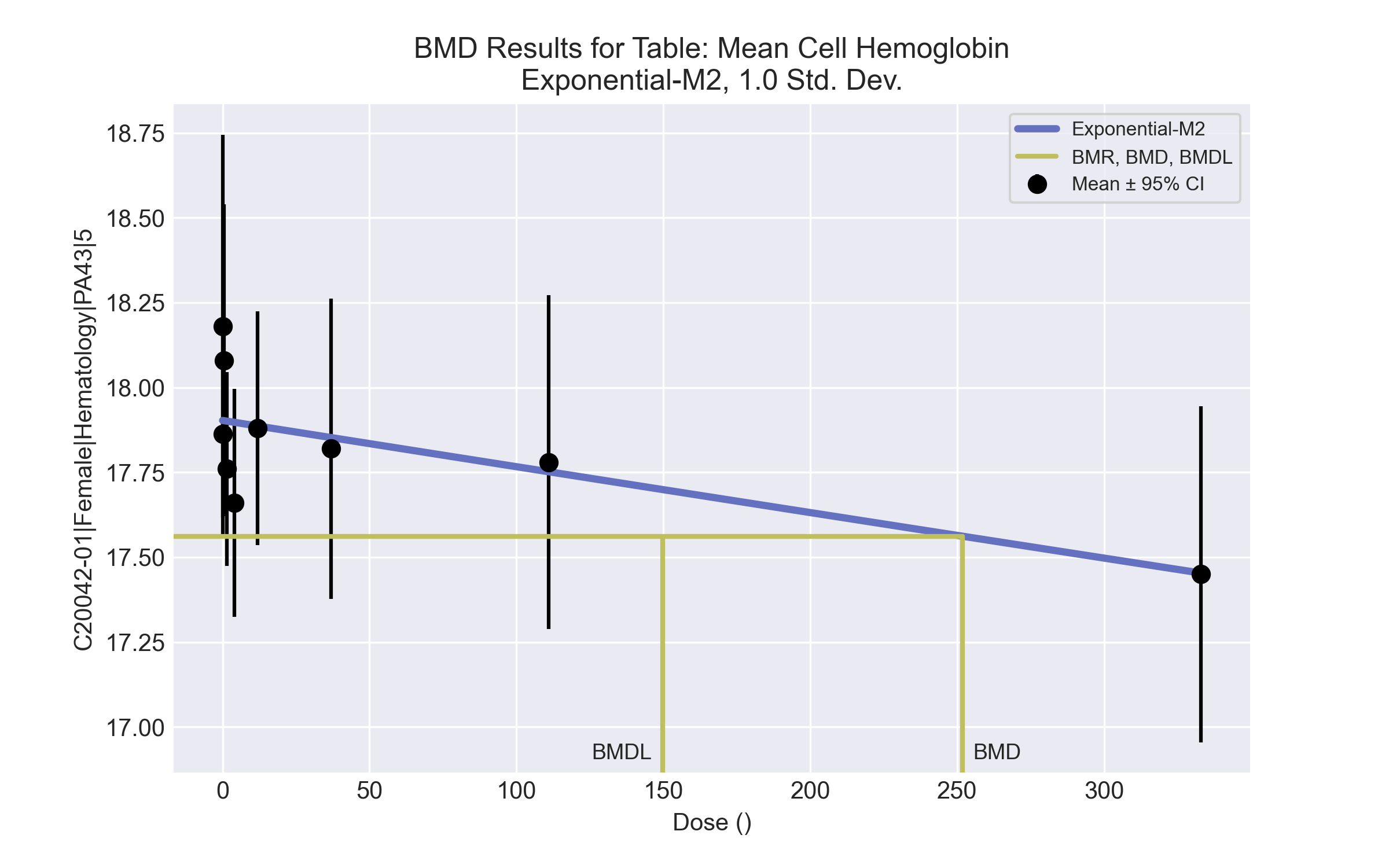
b Recommended model

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 2°) | Valid | - |
| Polynomial 3° | Valid | - |
| Polynomial 4° | Valid | - |
| Polynomial 5° | Valid | - |
| Polynomial 6° | Valid | - |
| Polynomial 7° | Valid | - |
| Polynomial 8° | Valid | - |
| Hill | Failure | **Failures**  • BMDL does not exist  **Cautions**  • Warning(s): BMDL computation failed. |
| Exponential M2a (equivalent models include Exponential M3) | Valid | - |
| Exponential M4 (equivalent models include Exponential M5) | Warning | **Warnings**  • BMD/BMDL ratio is greater than threshold (1.89e+02 > 20.0)  **Cautions**  • BMD/BMDL ratio is greater than threshold (1.89e+02 > 5.0) |

a Recommended model

## Recommended model



====================================================================   
 Exponential Model. (Version: 1.11; Date: 03/14/2017)   
 Input Data File: C:\Windows\TEMP\bmds-6yxynn\_g.(d)   
 Gnuplot Plotting File:   
 Tue Apr 13 13:58:42 2021  
 ====================================================================   
  
 BMDS\_Model\_Run   
~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~  
   
 The form of the response function by Model:   
 Model 2: Y[dose] = a \* exp{sign \* b \* dose}  
 Model 3: Y[dose] = a \* exp{sign \* (b \* dose)^d}  
 Model 4: Y[dose] = a \* [c-(c-1) \* exp{-b \* dose}]  
 Model 5: Y[dose] = a \* [c-(c-1) \* exp{-(b \* dose)^d}]  
  
 Note: Y[dose] is the median response for exposure = dose;  
 sign = +1 for increasing trend in data;  
 sign = -1 for decreasing trend.  
  
 Model 2 is nested within Models 3 and 4.  
 Model 3 is nested within Model 5.  
 Model 4 is nested within Model 5.  
  
  
 Dependent variable = Response  
 Independent variable = Dose  
 Data are assumed to be distributed: normally  
 Variance Model: exp(lnalpha +rho \*ln(Y[dose]))  
 rho is set to 0.  
 A constant variance model is fit.  
  
 Total number of dose groups = 9  
 Total number of records with missing values = 0  
 Maximum number of iterations = 500  
 Relative Function Convergence has been set to: 1e-008  
 Parameter Convergence has been set to: 1e-008  
  
 MLE solution provided: Exact  
  
  
 Initial Parameter Values  
  
 Variable Model 2  
 -------- --------  
 lnalpha -2.34324   
 rho 0 Specified  
 a 17.777   
 b 7.62082e-005   
 c 0 Specified  
 d 1 Specified  
  
  
  
 Parameter Estimates  
  
 Variable Model 2 Std. Err.  
 -------- ------- ---------  
 lnalpha -2.15287 0.02396  
 a 17.9028 0.0554374  
 b 7.63658e-005 3.02302e-005  
  
 NC = No Convergence  
  
  
 Table of Stats From Input Data  
  
 Dose N Obs Mean Obs Std Dev  
 ----- --- ---------- -------------  
 0 8 17.86 0.3583  
 0.15 5 18.18 0.455  
 0.5 5 18.08 0.3701  
 1.4 5 17.76 0.2302  
 4 5 17.66 0.2702  
 12 5 17.88 0.2775  
 37 5 17.82 0.3564  
 111 5 17.78 0.3962  
 333 4 17.45 0.3109  
  
  
 Estimated Values of Interest  
  
 Dose Est Mean Est Std Scaled Residual  
 ------ ---------- --------- ----------------  
 0 17.9 0.3408 -0.3341  
 0.15 17.9 0.3408 1.82  
 0.5 17.9 0.3408 1.167  
 1.4 17.9 0.3408 -0.9241  
 4 17.9 0.3408 -1.557  
 12 17.89 0.3408 -0.04172  
 37 17.85 0.3408 -0.2116  
 111 17.75 0.3408 0.186  
 333 17.45 0.3408 -0.01897  
  
  
  
 Other models for which likelihoods are calculated:  
  
 Model A1: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
  
 Model A2: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma(i)^2  
  
 Model A3: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = exp(lalpha + log(mean(i)) \* rho)  
  
 Model R: Yij = Mu + e(i)  
 Var{e(ij)} = Sigma^2  
  
  
 Likelihoods of Interest  
  
 Model Log(likelihood) DF AIC  
 ------- ----------------- ---- ------------  
 A1 31.56616 10 -43.13232  
 A2 33.38962 18 -30.77923  
 A3 31.56616 10 -43.13232  
 R 24.05161 2 -44.10323  
 2 27.09243 3 -48.18486  
  
  
 Additive constant for all log-likelihoods = -43.19. This constant added to the  
 above values gives the log-likelihood including the term that does not  
 depend on the model parameters.  
  
  
 Explanation of Tests  
  
 Test 1: Does response and/or variances differ among Dose levels? (A2 vs. R)  
 Test 2: Are Variances Homogeneous? (A2 vs. A1)  
 Test 3: Are variances adequately modeled? (A2 vs. A3)  
 Test 4: Does Model 2 fit the data? (A3 vs. 2)  
  
  
 Tests of Interest  
  
 Test -2\*log(Likelihood Ratio) D. F. p-value  
 -------- ------------------------ ------ --------------  
 Test 1 18.68 16 0.2858  
 Test 2 3.647 8 0.8875  
 Test 3 3.647 8 0.8875  
 Test 4 8.947 7 0.2565  
  
  
 The p-value for Test 1 is greater than .05. There may not be a  
 diffence between responses and/or variances among the dose levels  
 Modelling the data with a dose/response curve may not be appropriate.  
  
 The p-value for Test 2 is greater than .1. A homogeneous  
 variance model appears to be appropriate here.  
  
 The p-value for Test 3 is greater than .1. The modeled  
 variance appears to be appropriate here.  
  
 The p-value for Test 4 is greater than .1. Model 2 seems  
 to adequately describe the data.  
  
  
 Benchmark Dose Computations:  
  
 Specified Effect = 1.000000  
  
 Risk Type = Estimated standard deviations from control  
  
 Confidence Level = 0.950000  
  
 BMD = 251.686  
  
 BMDL = 149.752  
  
 BMDU = 762.475

# Female BMD Results for Table: Platelet count

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 8 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 4 | 5 |
| Mean ± SD | 967.5 ± 255.215 | 974.4 ± 161.769 | 984.6 ± 33.366 | 1054.8 ± 107.297 | 1048 ± 172.208 | 1150.4 ± 134.97 | 1197.2 ± 64.616 | 1012.8 ± 298.713 | 1123.5 ± 223.834 | 1183.6 ± 186.945 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°) | 0.034 | 601.584 | 1201 | 555.135 | No model was recommended as a best-fitting model. Doses were dropped until there were only 3 remaining dose-groups. |
| Polynomial 8° | 0.034 | 601.584 | 1200.87 | 555.135 |
| Hill | 0.063 | 600.865 | -999 | -999 |
| Exponential M2 (equivalent models include Exponential M3) | 0.034 | 601.608 | 1193.92 | 591.601 |
| Exponential M4 | 0.105 | 598.79 | -999 | 0 |
| Exponential M5 | 0.065 | 600.78 | -999 | 0 |

a Modeled variance case presented (BMDS Test 2 p-value = 0.001, BMDS Test 3 p-value = 0.011).

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.01076)  • Goodness of fit p-value is less than threshold (0.0338 < 0.1)  • BMD/high dose ratio is greater than threshold (1.2 > 1.0) |
| Polynomial 8° | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.01076)  • Goodness of fit p-value is less than threshold (0.0338 < 0.1)  • BMD/high dose ratio is greater than threshold (1.2 > 1.0) |
| Hill | Failure | **Failures**  • BMD does not exist  • BMDL does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.01076)  • Goodness of fit p-value is less than threshold (0.0633 < 0.1)  **Cautions**  • Warning(s): BMR value is not in the range of the mean function |
| Exponential M2 (equivalent models include Exponential M3) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.01076)  • Goodness of fit p-value is less than threshold (0.0336 < 0.1)  • BMD/high dose ratio is greater than threshold (1.19 > 1.0) |
| Exponential M4 | Failure | **Failures**  • BMD does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.01076) |
| Exponential M5 | Failure | **Failures**  • BMD does not exist  **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.01076)  • Goodness of fit p-value is less than threshold (0.0653 < 0.1) |

## Recommended model

*No model was recommended as a best-fitting model.*

# Female BMD Results for Table: Reticulocyte count

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37 | 111 | 333 | 1000 |
| N | 8 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 4 | 5 |
| Mean ± SD | 227.162 ± 44.806 | 254.48 ± 27.037 | 210.6 ± 34.657 | 210.76 ± 55.873 | 225.44 ± 37.913 | 209.22 ± 8.416 | 188 ± 25.592 | 177.26 ± 24.292 | 134.525 ± 31.921 | 138.84 ± 35.493 |

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 3°, 5°) | 0.005 | 442.136 | 434.65 | 316.444 | No model was recommended as a best-fitting model. Doses were dropped until there were only 3 remaining dose-groups. |
| Polynomial 2° | 0.005 | 442.136 | 434.649 | 273.238 |
| Polynomial 4° | 0.005 | 442.136 | 434.648 | 245.823 |
| Polynomial 6° | <0.0001 | 970.837 | -9999 | 189.37 |
| Polynomial 7° | <0.0001 | 457.842 | -9999 | 745.272 |
| Polynomial 8° | <0.0001 | 459.842 | -9999 | 190.738 |
| Hill | 0.548 | 427.916 | 36.248 | -999 |
| Exponential M2 (equivalent models include Exponential M3) | 0.01 | 439.972 | 299.305 | 110.651 |
| Exponential M4 (equivalent models include Exponential M5) | 0.515 | 428.201 | 52.664 | 23.346 |

a Modeled variance case presented (BMDS Test 2 p-value = 0.051, BMDS Test 3 p-value = 0.039).

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 3°, 5°) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.03889)  • Goodness of fit p-value is less than threshold (0.00464 < 0.1) |
| Polynomial 2° | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.03889)  • Goodness of fit p-value is less than threshold (0.00464 < 0.1) |
| Polynomial 4° | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.03889)  • Goodness of fit p-value is less than threshold (0.00464 < 0.1) |
| Polynomial 6° | Warning | **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.03889)  • Residual at lowest dose is greater than threshold (3.05 > 2.0)  • Ratio of modeled to actual stdev. at control is greater than threshold (11.0 > 1.5)  **Cautions**  • Warning(s): THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!; BMD = 100\*(maximum dose) |
| Polynomial 7° | Warning | **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.03889)  **Cautions**  • Warning(s): BMD = 100\*(maximum dose) |
| Polynomial 8° | Warning | **Warnings**  • Residual of Interest does not exist  • Variance model poorly fits dataset (p-value 3 = 0.03889)  **Cautions**  • Warning(s): BMD = 100\*(maximum dose) |
| Hill | Failure | **Failures**  • BMDL does not exist  **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.03889)  **Cautions**  • Warning(s): BMDL computation failed. |
| Exponential M2 (equivalent models include Exponential M3) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.03889)  • Goodness of fit p-value is less than threshold (0.0104 < 0.1) |
| Exponential M4 (equivalent models include Exponential M5) | Warning | **Warnings**  • Variance model poorly fits dataset (p-value 3 = 0.03889) |

## Recommended model

*No model was recommended as a best-fitting model.*

# Female BMD Results for Table: Thyroid Stimulating Hormone

BMDS version: BMDS v2.7.0

## Input dataset

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Dose | 0 | 0.15 | 0.5 | 1.4 | 4 | 12 | 37a | 111a | 333a | 1000a |
| N | 6 | 5 | 5 | 5 | 4 | 5 | 4 | 5 | 4 | 5 |
| Mean ± SD | 2.783 ± 0.799 | 3.36 ± 1.907 | 2.94 ± 1.537 | 4.2 ± 1.065 | 4.025 ± 0.665 | 4.26 ± 1.101 | 4.475 ± 0.479 | 5.48 ± 2.463 | 4.9 ± 2.719 | 8.6 ± 3.941 |

a Dose group removed in BMD modeling session

## Summary table

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| Modela | Goodness of fit | | BMD | BMDL | Comments |
| *p*-value | AIC |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | 0.353 | 47.436 | 12.541 | 6.575 | Hill recommended as best-fitting model on the basis of the lowest AIC. |
| Hillb | 0.84 | 45.86 | 1.078 | 0.267 |
| Exponential M2 (equivalent models include Exponential M3) | 0.335 | 47.584 | 12.973 | 7.529 |
| Exponential M4 | 0.643 | 46.693 | 1.801 | 0.251 |
| Exponential M5 | 0.658 | 47.86 | 1.281 | 0.512 |

a Constant variance case presented (BMDS Test 2 p-value = 0.197, BMDS Test 3 p-value = 0.197).

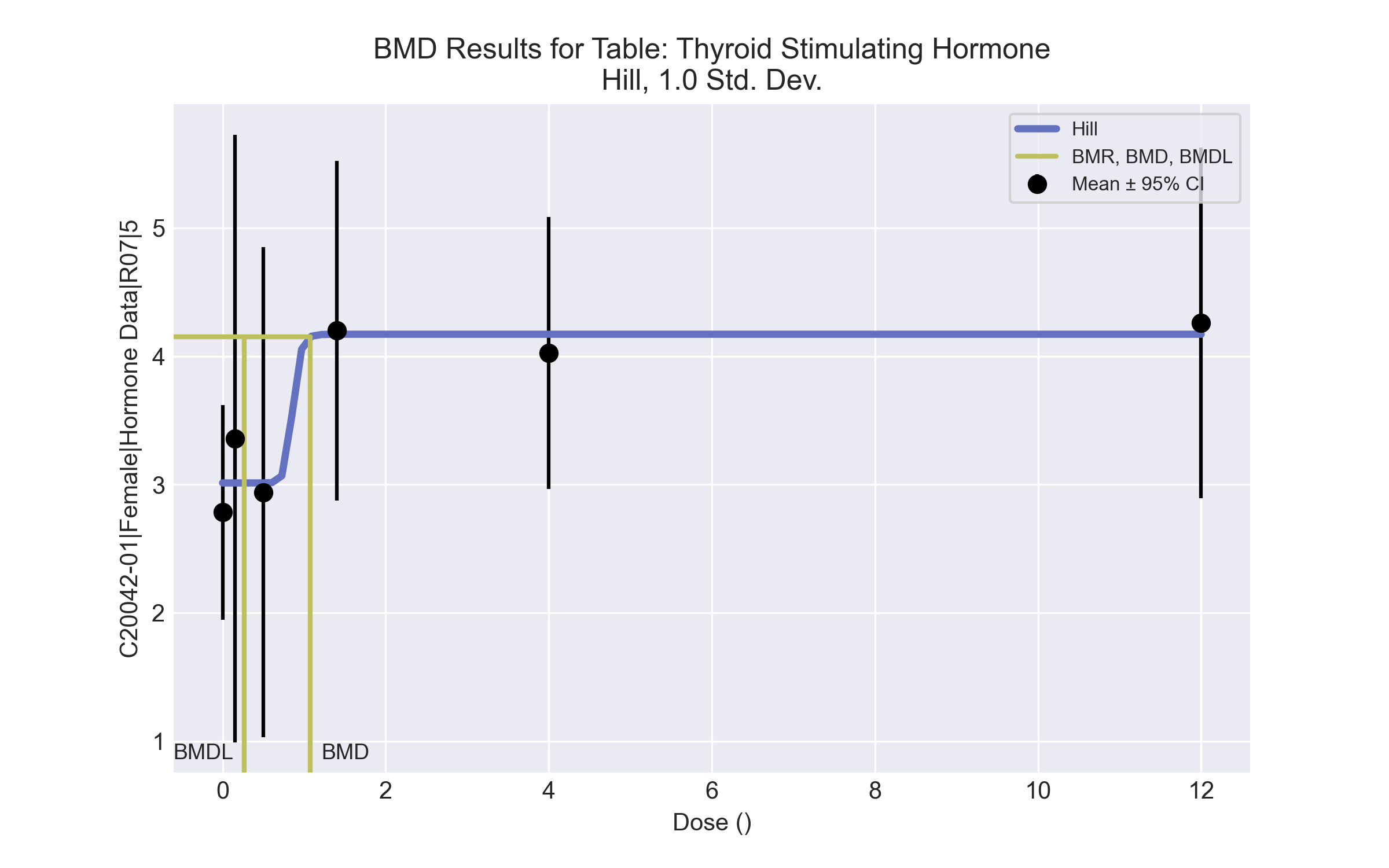
b Recommended model

## Model recommendation details

|  |  |  |
| --- | --- | --- |
| Model | Bin | Notes |
| Linear (equivalent models include Power, Polynomial 2°, 3°, 4°, 5°, 6°, 7°, 8°) | Warning | **Warnings**  • BMD/high dose ratio is greater than threshold (1.05 > 1.0)  • Ratio of modeled to actual stdev. at control is greater than threshold (1.51 > 1.5) |
| Hilla | Valid | - |
| Exponential M2 (equivalent models include Exponential M3) | Warning | **Warnings**  • BMD/high dose ratio is greater than threshold (1.08 > 1.0) |
| Exponential M4 | Valid | **Cautions**  • BMD/BMDL ratio is greater than threshold (7.19 > 5.0) |
| Exponential M5 | Valid | - |

a Recommended model

## Recommended model



====================================================================   
 Hill Model. (Version: 2.18; Date: 03/14/2017)   
 Input Data File: C:\Windows\TEMP\bmds-ae61de6y.(d)   
 Gnuplot Plotting File: C:\Windows\TEMP\bmds-ae61de6y.plt  
 Tue Apr 13 14:02:33 2021  
 ====================================================================   
  
 BMDS\_Model\_Run   
~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~  
   
 The form of the response function is:   
  
 Y[dose] = intercept + v\*dose^n/(k^n + dose^n)  
  
  
 Dependent variable = Response  
 Independent variable = Dose  
 rho is set to 0  
 Power parameter restricted to be greater than 1  
 A constant variance model is fit  
  
 Total number of dose groups = 6  
 Total number of records with missing values = 0  
 Maximum number of iterations = 500  
 Relative Function Convergence has been set to: 1e-008  
 Parameter Convergence has been set to: 1e-008  
  
  
  
 Default Initial Parameter Values   
 alpha = 1.57966  
 rho = 0 Specified  
 intercept = 2.78333  
 v = 1.47667  
 n = 0.998488  
 k = 1.88452  
  
  
 Asymptotic Correlation Matrix of Parameter Estimates  
  
 ( \*\*\* The model parameter(s) -rho -n   
 have been estimated at a boundary point, or have been specified by the user,  
 and do not appear in the correlation matrix )  
  
 alpha intercept v k  
  
 alpha 1 -8.3e-008 1.9e-007 -2.3e-007  
  
 intercept -8.3e-008 1 -0.68 0.012  
  
 v 1.9e-007 -0.68 1 0.016  
  
 k -2.3e-007 0.012 0.016 1  
  
  
  
 Parameter Estimates  
  
 95.0% Wald Confidence Interval  
 Variable Estimate Std. Err. Lower Conf. Limit Upper Conf. Limit  
 alpha 1.29954 0.33554 0.641894 1.95718  
 intercept 3.01248 0.28502 2.45385 3.57111  
 v 1.15901 0.417269 0.341175 1.97684  
 n 18 NA  
 k 0.858502 7.68332 -14.2005 15.9175  
  
NA - Indicates that this parameter has hit a bound  
 implied by some inequality constraint and thus  
 has no standard error.  
  
  
  
 Table of Data and Estimated Values of Interest  
  
 Dose N Obs Mean Est Mean Obs Std Dev Est Std Dev Scaled Res.  
------ --- -------- -------- ----------- ----------- ----------  
  
 0 6 2.78 3.01 0.799 1.14 -0.492  
 0.15 5 3.36 3.01 1.91 1.14 0.682  
 0.5 5 2.94 3.01 1.54 1.14 -0.142  
 1.4 5 4.2 4.17 1.07 1.14 0.0563  
 4 4 4.03 4.17 0.665 1.14 -0.257  
 12 5 4.26 4.17 1.1 1.14 0.174  
  
  
  
 Model Descriptions for likelihoods calculated  
  
  
 Model A1: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
  
 Model A2: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma(i)^2  
  
 Model A3: Yij = Mu(i) + e(ij)  
 Var{e(ij)} = Sigma^2  
 Model A3 uses any fixed variance parameters that  
 were specified by the user  
  
 Model R: Yi = Mu + e(i)  
 Var{e(i)} = Sigma^2  
  
  
 Likelihoods of Interest  
  
 Model Log(likelihood) # Param's AIC  
 A1 -18.510987 7 51.021974  
 A2 -14.843525 12 53.687051  
 A3 -18.510987 7 51.021974  
 fitted -18.930143 4 45.860286  
 R -22.363833 2 48.727665  
  
  
 Explanation of Tests   
  
 Test 1: Do responses and/or variances differ among Dose levels?   
 (A2 vs. R)  
 Test 2: Are Variances Homogeneous? (A1 vs A2)  
 Test 3: Are variances adequately modeled? (A2 vs. A3)  
 Test 4: Does the Model for the Mean Fit? (A3 vs. fitted)  
 (Note: When rho=0 the results of Test 3 and Test 2 will be the same.)  
  
 Tests of Interest   
  
 Test -2\*log(Likelihood Ratio) Test df p-value   
  
 Test 1 15.0406 10 0.1306  
 Test 2 7.33492 5 0.1969  
 Test 3 7.33492 5 0.1969  
 Test 4 0.838312 3 0.8403  
  
The p-value for Test 1 is greater than .05. There may not be a  
diffence between responses and/or variances among the dose levels  
Modelling the data with a dose/response curve may not be appropriate  
  
The p-value for Test 2 is greater than .1. A homogeneous variance   
model appears to be appropriate here  
  
  
The p-value for Test 3 is greater than .1. The modeled variance appears   
 to be appropriate here  
  
The p-value for Test 4 is greater than .1. The model chosen seems   
to adequately describe the data  
   
  
 Benchmark Dose Computation  
  
Specified effect = 1  
  
Risk Type = Estimated standard deviations from the control mean   
  
Confidence level = 0.95  
  
 BMD = 1.07767  
  
 BMDL = 0.266926  
  
 BMDU = 1.44e+006