

Annex to:

EFSA CONTAM Panel (EFSA Panel on Contaminants in the Food Chain), Alexander J, Barregård L, Bignami M, Brüschweiler B, Ceccatelli S, Cottrill B, Dinovi M, Grasl-Kraupp B, Hogstrand C, Hoogenboom LR, Knutsen HK, Nebbia CS, Oswald IP, Petersen A, Rose M, Roudot A-C, Schwerdtle T, Vleminckx C, Vollmer G, Wallace H, De Saeger S, Sundstøl Eriksen G, Farmer P, Fremy J-M, Gong YY, Meyer K, Naegeli H, Parent-Massin D, Rietjens I, van Egmond H, Altieri A, Eskola M, Gergelova P, Ramos Bordajandi L, Benkova B, Dörr B, Gkrillas A, Gustavsson N, van Manen M and Edler L, 2017. Scientific opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed. *EFSA Journal* 2017;15(9):4718, 345 pp. doi:10.2903/j.efsa.2017.4718

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Annex A – Supplementary information for the BMD calculations

Supplementary information IVERSONBW (Iverson et al., 1995)

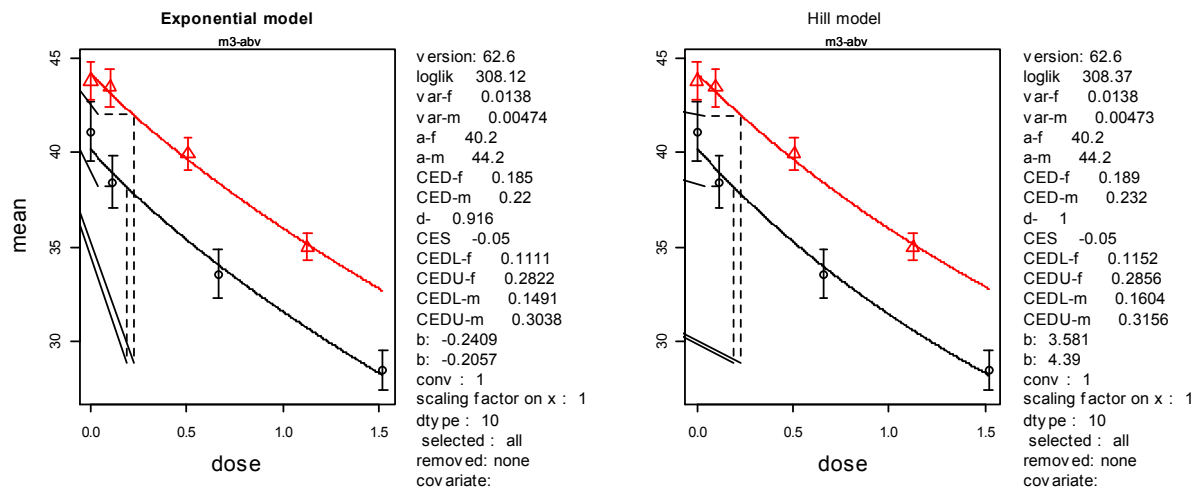
Details of the BMD Analysis of the body weight data of Iverson et al. (1995) reported in Appendix G, Section G.2.1 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

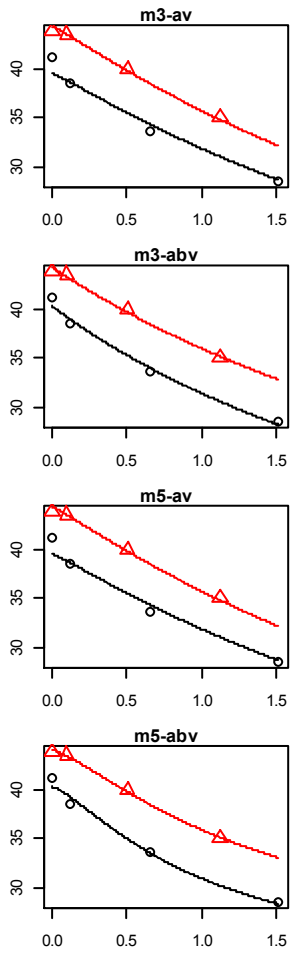
Filename: IVERSONBW Folder IVERSON-1995-mice

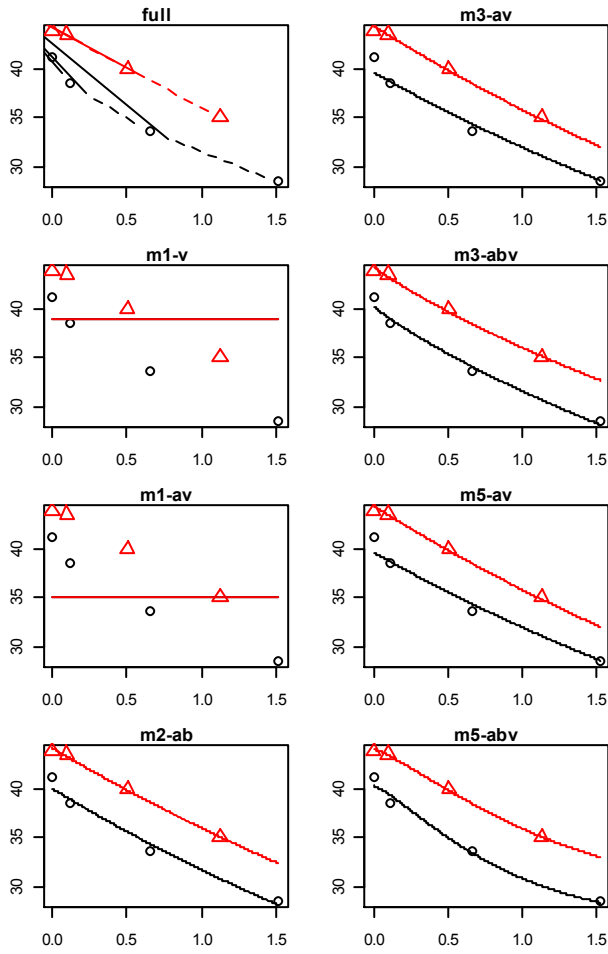
Output from PROAST 62.6 from 13 OCT 2016

BMD analysis of body weight data of females (f: black curve) and males (m: red curve) combined

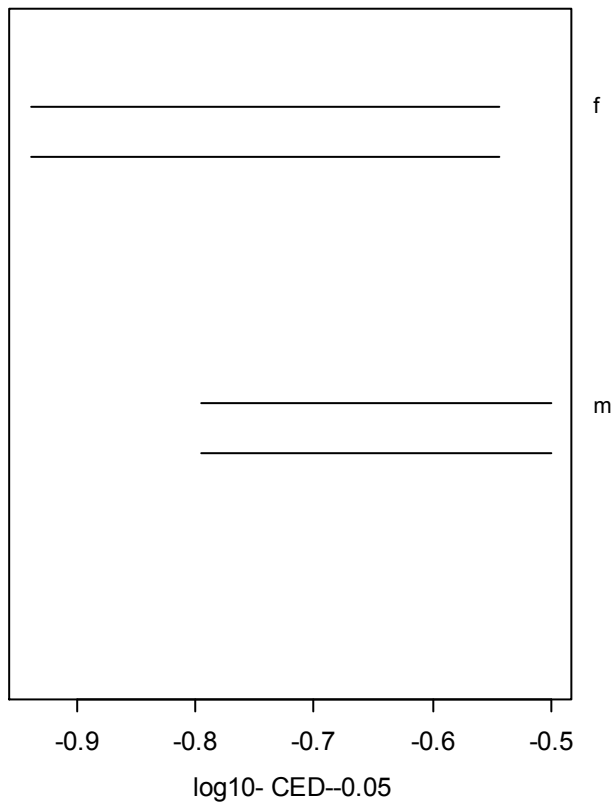
Graphical output







Graphic of the BMD- 90% confidence interval (C.I.) for female and male and for exponential and Hill models



Numerical output

Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	9	289.36	-560.72
full-v	1	10	310.36	-600.72
m1-v	1	3	135.59	-265.18
m1-av	1	4	162.95	-317.9
m2-ab	1	5	285.98	-561.96
m3-av	1	6	306.94	-601.88
m3-abv	1	7	308.12	-602.24
m5-av	1	7	306.94	-599.88
m5-abv	1	8	308.95	-601.9

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-abv.

However, m5-abv is a reasonable model as well, and so is m3-av.

Selected model: m3-abv

The CED – Critical Effect Dose (in original units) and the 90 % C.I. for group f is:

0.18484

0.1111

0.28222

The CED (in original units) and the 90 % C.I. for group m is:
 0.21964
 0.14911
 0.30377

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-av	1	6	306.86	-601.72
m3-abv	1	7	308.37	-602.74
m5-av	1	7	306.86	-599.72
m5-abv	1	8	308.82	-601.64

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-abv.
 However, m3-av is a reasonable model as well, and so is m5-abv.

Selected model: m3-abv

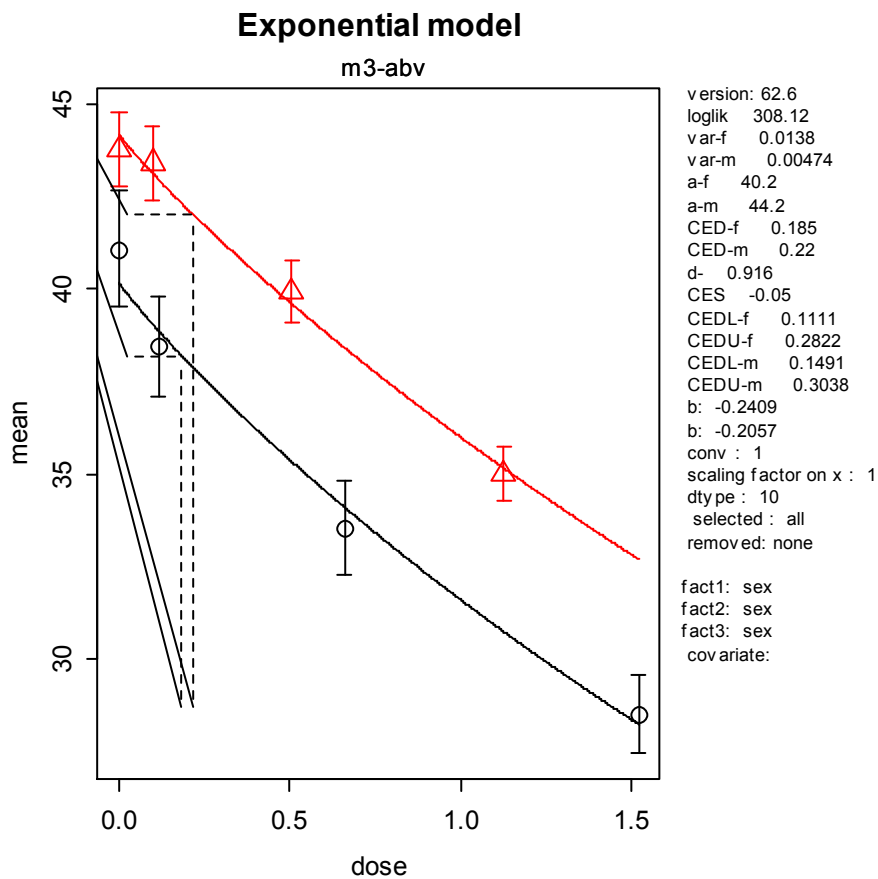
The CED (in original units) and the 90 % C.I. for group f is:
 0.18929
 0.11516
 0.28557

The CED (in original units) and the 90 % C.I. for group m is:
 0.23205
 0.16044
 0.31558

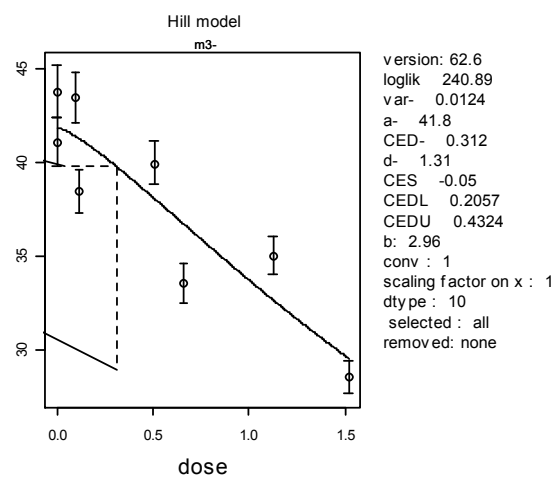
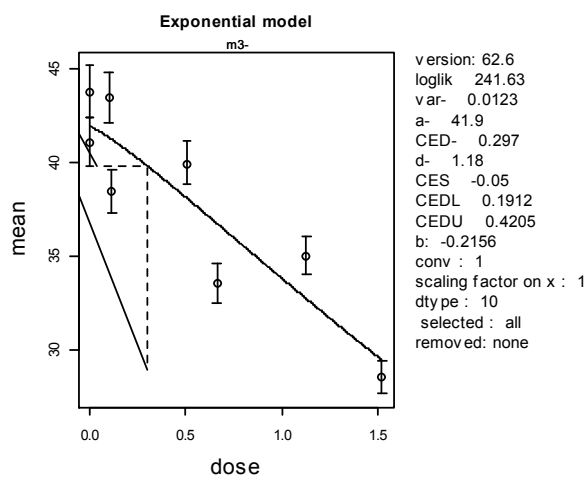
covar	LB	UB	LB.H	UB.H
1 f	0.11516	0.28557	0.11516	0.28557
2 m	0.16044	0.31558	0.16044	0.31558

f: female; m: male; LB: lower bound; UB: upper bound;

Model with the curve (black for females) showing the smallest BMDL of 0.1111



BMD analysis of the pooled data of females and males with smallest BMDL of 0.1912



File of Iverson data: IVERSONBW

```
5
0      0      10      0      0
dose  n      mean  sd     sex
0     37     43.85  2.69  m
0.098 35     43.51  2.86  m
0.506 43     40.03  3.00  m
1.126 42     35.09  2.56  m
0     36     41.54  6.26  f
0.115 42     38.71  4.73  f
0.661 37     33.76  3.92  f
1.520 38     28.55  2.08  f
```

Supplementary information IVERSONFDC (Iverson et al., 1995):

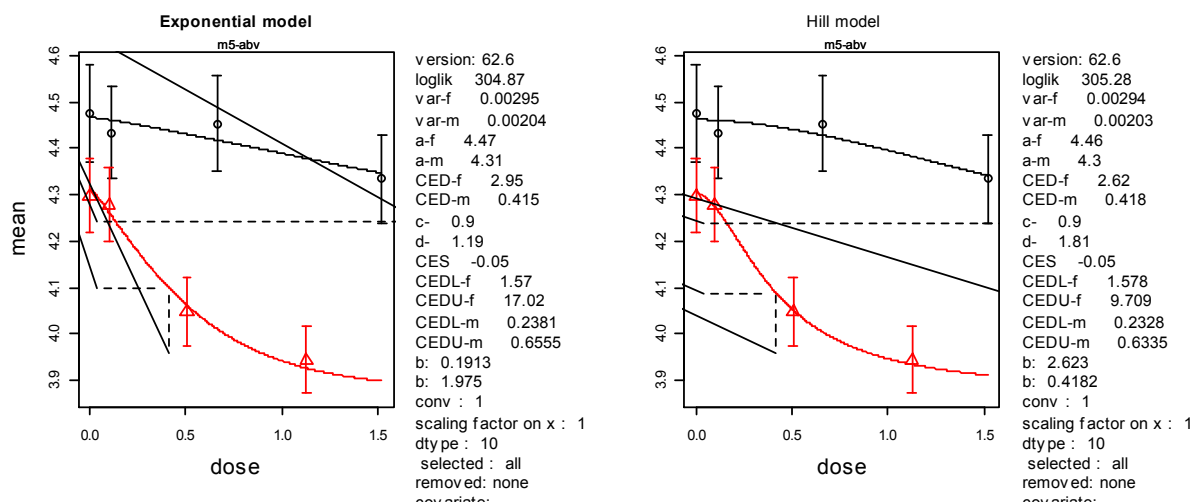
Details of the BMD Analysis of the feed intake data of Iverson et al. (1995) reported in Appendix G, Section G.2.1 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

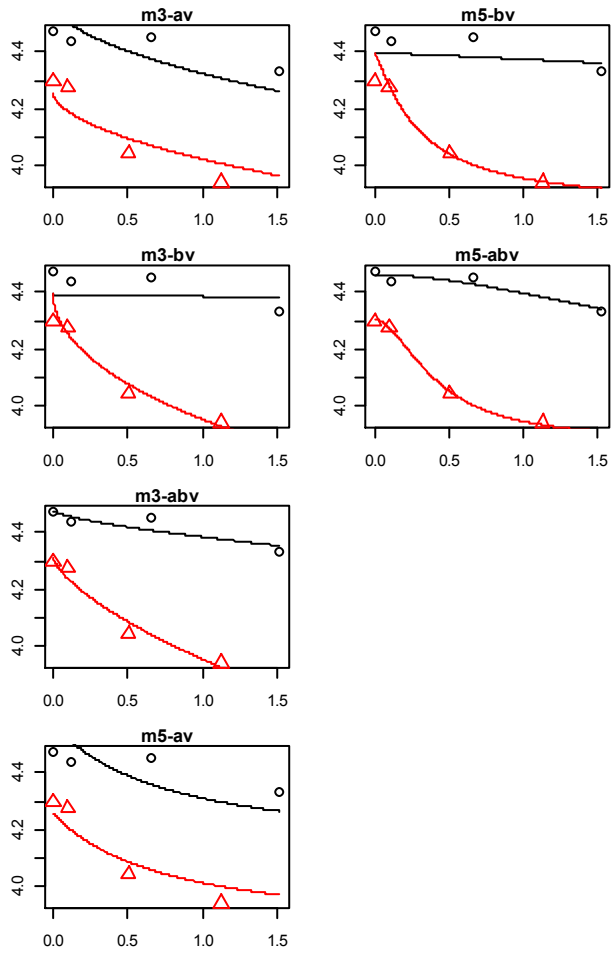
Filename: IVERSONFDC Folder IVERSON-1995-mice

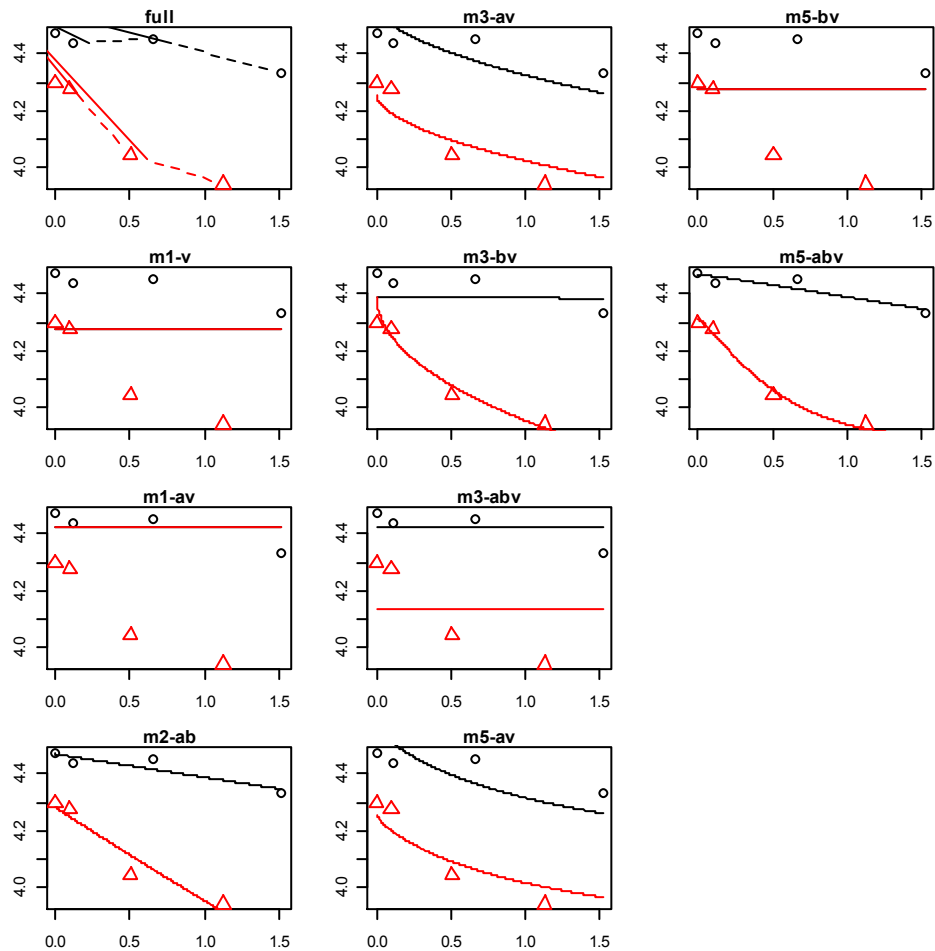
Output from PROAST 62.6 from 13 OCT 2016

BMD analysis of feed intake data of females (f: black curve) and males (m: red curve) combined

Graphical output







Numerical output

Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	9	304.03	-590.06
full-v	1	10	305.63	-591.26
m1-v	1	3	249.86	-493.72
m1-av	1	4	278.41	-548.82
m2-ab	1	5	301.65	-593.3
m3-av	1	6	295.5	-579
m3-bv	1	6	299.27	-586.54
m3-abv	1	7	278.41	-542.82
m5-av	1	7	295.96	-577.92
m5-bv	1	7	249.86	-485.72
m5-abv	1	8	304.87	-593.74

AIC: Akaike's Information Criterion.

Best model with covariates is: m5-abv.
However, m2-ab is a reasonable model as well.

Selected model: m5-abv

The CED (in original units) and the 90 % C.I. for group f is:
 2.9472
 1.5698
 17.023

The CED (in orig. units) and the 90 % C.I. for group m is:
 0.41508
 0.23807
 0.65547

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-av	1	6	295.53	-579.06
m3-bv	1	6	299.31	-586.62
m3-abv	1	7	303.85	-593.7
m5-av	1	7	296.24	-578.48
m5-bv	1	7	300.08	-586.16
m5-abv	1	8	305.28	-594.56

AIC: Akaike's Information Criterion.

Best model with covariates is: m5-abv.
 However, m3-abv is a reasonable model as well.

Selected model: m5-abv

The CED (in original units) and the 90 % C.I. for group f is:
 2.6228
 1.5775
 9.7092

The CED (in original units) and the 90 % C.I. for group m is:
 0.4182
 0.23281
 0.63349

File of Iverson data: IVERSONFDC

```

5
0    0    10    0    0
dose n    mean sd   sex
0    24    4.30 0.16  m
0.098 24    4.28 0.17  m
0.506 25    4.05 0.17  m
1.126 25    3.95 0.24  m
0    22    4.48 0.25  f
0.115 24    4.44 0.23  f
0.661 23    4.46 0.26  f
1.520 25    4.34 0.24  f

```

Supplementary information IVERSON-concentration–response (Iverson et al., 1995)

Details of the BMD Analysis of the body weight data of Iverson et al. (1995) reported in Appendix G, Section G.2.2 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed - based on the concentration of DON. The data file is not shown below.

Folder IVERSON-1995-mice

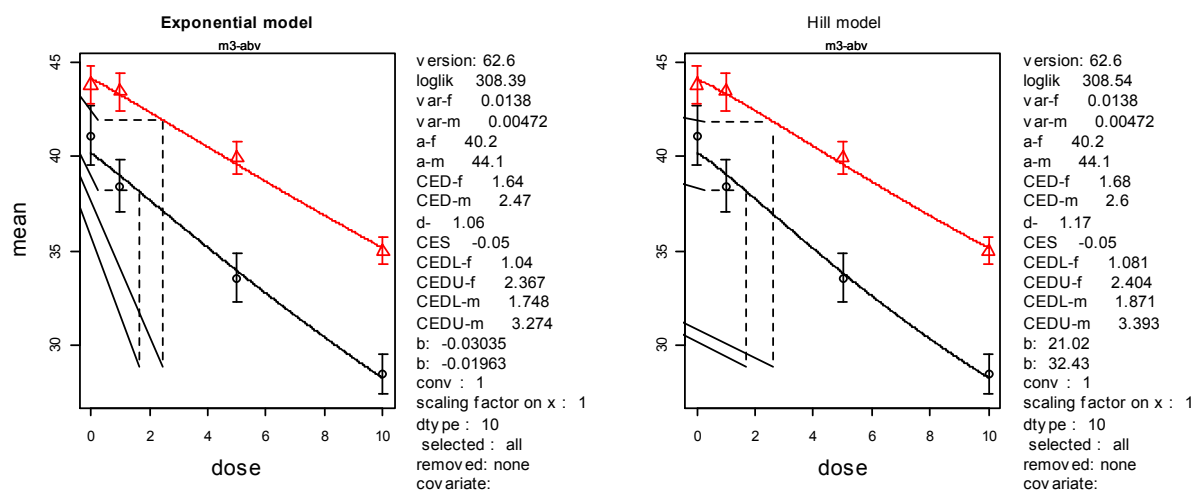
Output from PROAST 62.6 from 13 OCT 2016

BMD analysis of **body weight data** of females (f: black curve) and males (m: red curve) combined:

Concentration-response analyses

Graphical output

body weight



Numerical output

Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	9	289.36	-560.72
full-v	1	10	310.36	-600.72
m1-v	1	3	135.59	-265.18
m1-av	1	4	162.95	-317.9
m2-ab	1	5	287.47	-564.94
m3-av	1	6	298.84	-585.68
m3-abv	1	7	308.39	-602.78
m5-av	1	7	298.84	-583.68
m5-abv	1	8	308.8	-601.6

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-abv.

However, m5-abv is a reasonable model as well.

Selected model: m3-abv

The CED (in orig. units) and the 90 % C.I. for group f is:
 1.6374
 1.0404
 2.3674

The CED (in orig. units) and the 90 % C.I. for group m is:
 2.4671
 1.7484
 3.2742

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-av	1	6	298.81	-585.62
m3-abv	1	7	308.54	-603.08
m5-av	1	7	298.81	-583.62
m5-abv	1	8	308.7	-601.4

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-abv.
 However, m5-abv is a reasonable model as well.

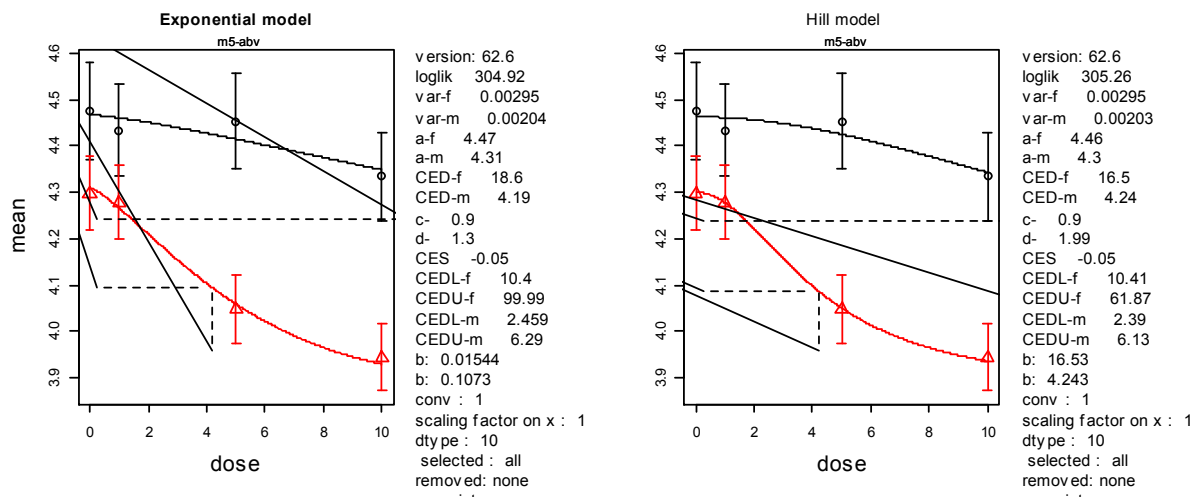
Selected model: m3-abv

The CED (in original units) and the 90 % C.I. for group f is:
 1.6831
 1.0813
 2.4044

The CED (in original units) and the 90 % C.I. for group m is:
 2.5968
 1.8713
 3.3931

BMD analysis of **feed intake data** of females (f: black curve) and males (m: red curve) combined:
 Concentration-response analyses

Graphical output



Numerical output

Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	9	304.03	-590.06
full-v	1	10	305.63	-591.26
m1-v	1	3	249.86	-493.72
m1-av	1	4	278.41	-548.82
m2-ab	1	5	302.14	-594.28
m3-av	1	6	298.01	-584.02
m3-bv	1	6	299.47	-586.94
m3-abv	1	7	278.41	-542.82
m5-av	1	7	298.1	-582.2
m5-bv	1	7	249.86	-485.72
m5-abv	1	8	304.92	-593.84

AIC: Akaike's Information Criterion.

Best model with covariates is: m2-ab.
However, m5-abv is a reasonable model as well.

Selected model: m5-abv

The CED (in original units) and the 90 % C.I. for group f is:

18.589
10.397
99.99

The CED (in original units) and the 90 % C.I. for group m is:

4.1924
2.4587
6.2895

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-av	1	6	298.02	-584.04
m3-bv	1	6	299.5	-587
m3-abv	1	7	304.05	-594.1
m5-av	1	7	298.11	-582.22
m5-bv	1	7	300	-586
m5-abv	1	8	305.26	-594.52

AIC: Akaike's Information Criterion.

Best model with covariates is: m5-abv.
However, m3-abv is a reasonable model as well.

Selected model: m5-abv

The CED (in original units) and the 90 % C.I. for group f is:

16.535
10.409
61.874

The CED (in original units) and the 90 % C.I. for group m is:

4.2431

2.3905

6.1303

Supplementary information BONDY (Bondy et al., 2016)

Details of the BMD Analysis of the body weight data of Bondy et al. (2016) reported in Appendix G, Section G.2.3 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

Filename: bondy Folder bondy

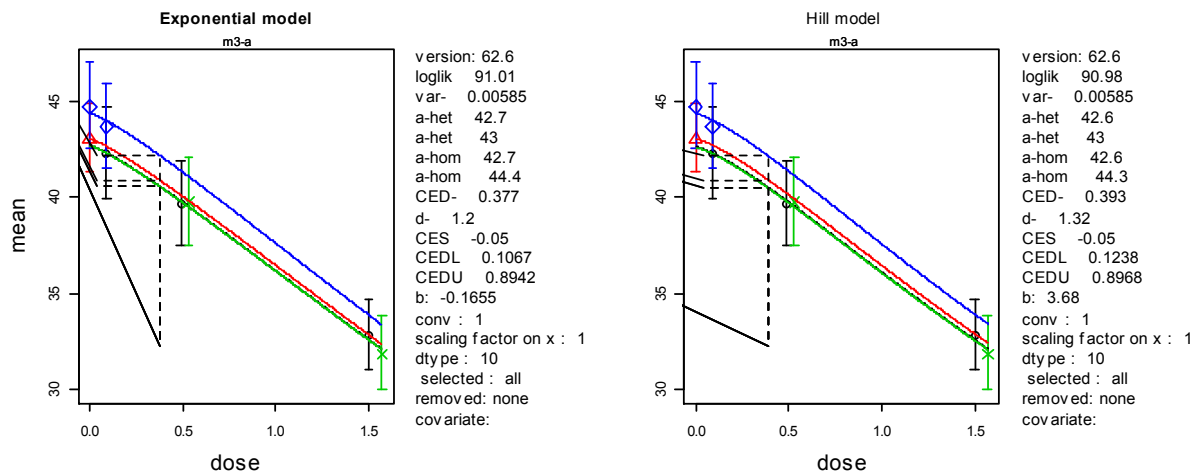
The dose-response evaluation used the BMD approach for the body weight data reported as means and standard deviations (sd) of the two experiments on transgenic mice of type p53+/+ (hom) and p53+/- (het) genotypes (type), combined, as shown in the table below. The number of mice per dose group (n) was 10 except for the highest dose of the p53+/+ type.

BONDYBW				
dose	n	mean	sd	type
mg/kg				
bw				
per day				
0	10	44.8	2.9	hom
0.09	10	43.8	3.7	hom
0.53	10	39.9	3.4	hom
1.57	9	32.0	2.8	hom
0	10	43.1	2.5	het
0.09	10	42.5	4.9	het
0.49	10	39.7	2.6	het
1.5	10	32.9	2.3	het

Data set used for PROAST 62.6 from 31 OCT 2016

Data set name: BONDYBW

Graphical output



Numerical output

Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	9	91.24	-164.48
full-v	1	12	92.44	-160.88
m1-	1	2	42.26	-80.52
m1-a	1	5	59.09	-108.18
m2-ab	1	9	91.2	-164.4
m3-a	1	7	91.01	-168.02
m3-b	1	7	90.22	-166.44
m3-ab	1	10	91.24	-162.48
m5-a	1	8	91.01	-166.02
m5-b	1	8	90.22	-164.44
m5-ab	1	11	91.24	-160.48

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-a.

However, m3-b is a reasonable model as well, and so is m5-a.

Selected model: m3-a

The CED (in original units) and the 90 % C.I. is:

0.37725

0.1067

0.89421

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-a	1	7	90.98	-167.96
m3-b	0	7	90.16	-166.32
m3-ab	1	10	91.24	-162.48
m5-a	1	8	90.98	-165.96
m5-b	1	8	90.16	-164.32
m5-ab	1	11	91.24	-160.48

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-a.

However, m3-b is a reasonable model as well, and so is m5-a.

Selected model: m3-a

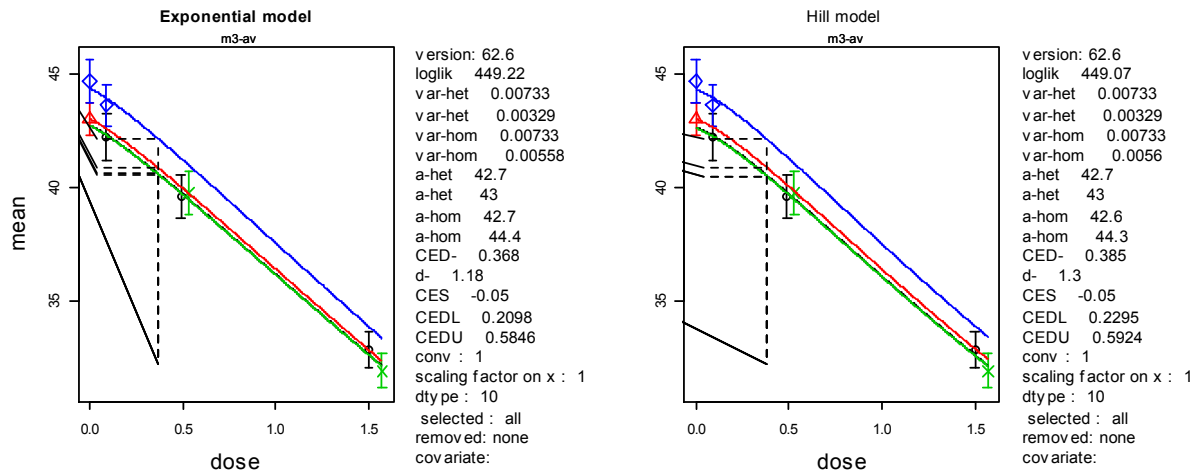
The CED (in original units) and the 90 % C.I. is:

0.39322

0.12379

0.89682

Tentative BMD Analysis of the data of Bondy et al. (2016) assuming the same doses and responses in a design with N=400 animals



Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	9	444.14	-870.28
full-v	1	12	450.24	-876.48
m1-v	1	5	230.71	-451.42
m1-av	1	8	329.12	-642.24
m2-ab	1	9	443.99	-869.98
m3-av	1	10	449.22	-878.44
m3-bv	1	10	444.89	-869.78
m3-abv	1	13	378.3	-730.6
m5-av	1	11	449.22	-876.44
m5-bv	0	11	234.21	-446.42
m5-abv	1	14	450.24	-872.48

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-av.
However, m5-av is a reasonable model as well.

Selected model: m3-av

Calculating the confidence interval (C.I.)

The CED (in original units) and the 90 % C.I. is:
0.36775
0.20978
0.58456

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-av	1	10	449.07	-878.14
m3-bv	1	10	444.64	-869.28
m3-abv	1	13	450.24	-874.48
m5-av	1	11	449.07	-876.14
m5-bv	1	11	444.64	-867.28
m5-abv	1	14	450.24	-872.48

AIC: Akaike's Information Criterion.

Best model with covariates is: m3-av.
However, m5-av is a reasonable model as well.

Selected model: m3-av

Calculating the confidence interval (C.I.)

The CED (in original units) and the 90 % C.I. is:
0.38496
0.22948
0.5924

Supplementary information SPRANDO2005ALL (Sprando et al., 2005)

Details of the BMD Analysis of the body weight data of Sprando et al. (2005) reported in Appendix G, Section G.3.1 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed - in male SD rats

Filename: Sprando2005all Folder Sprando-2005

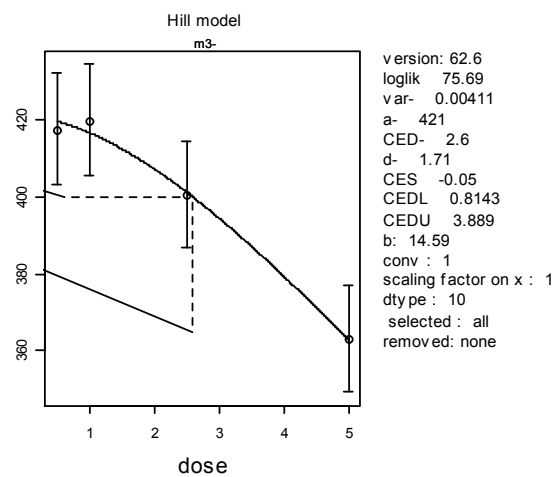
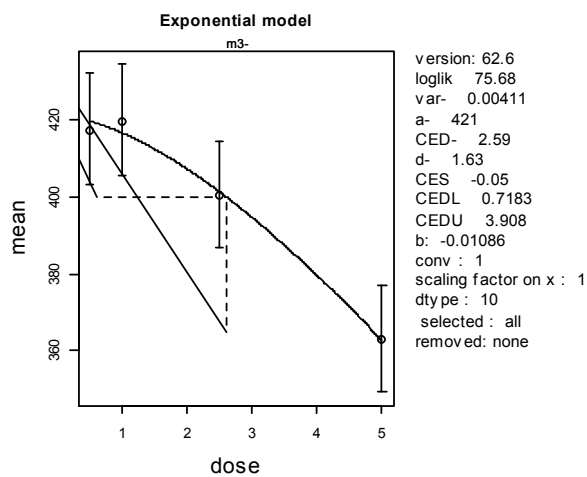
Output from PROAST 62.6 from 26 OCT 2016

Data, graphic and numerical output for body weight

SprandoBBWbis29

```

5
0 0 10 0 0
dose n mean sem sex
0 15 420.5 7.0 m
0.5 15 418.8 9.0 m
1 15 420.6 7.1 m
2.5 15 400.9 6.3 m
5 12 363.3 5.0 m
    
```



Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	5	75.85	-141.7
m1-	1	2	59.9	-115.8
m2-	1	3	75.16	-144.32
m3-	1	4	75.68	-143.36
m5-	1	5	75.82	-141.64

AIC: Akaike's Information Criterion.

Selected model: m3-

The CED (in original units) and the 90 % C.I. is:

- 2.5946
- 0.7183
- 3.9081

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-	1	4	75.69	-143.38
m5-	1	5	75.82	-141.64

AIC: Akaike’s Information Criterion.

Selected model: m3-

The CED (in original units) and the 90 % C.I. is:

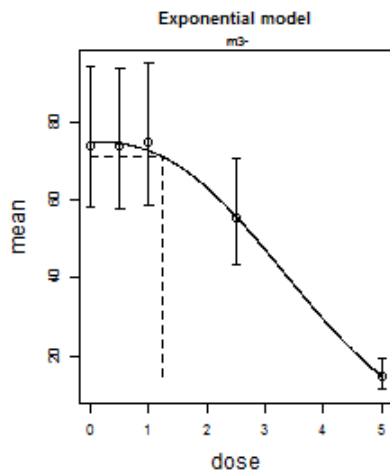
2.6033
0.81428
3.8893

Data, graphic and numerical output for average body weight gain by day 29

SprandoBWbis29

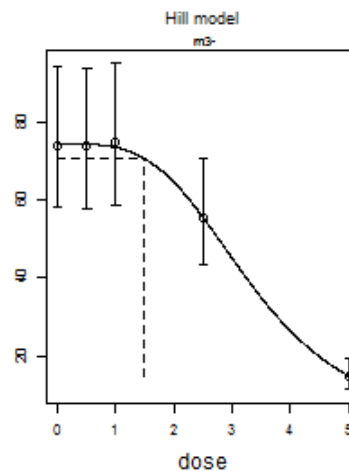
```

5
0 0 10 0 0
dose n mean sem sex
0 15 77.4 6.1 m
0.5 15 77.3 6.2 m
1 15 78.5 6.5 m
2.5 15 59.5 6.0 m
5 12 22.1 7.1 m
    
```



```

version: 62.6
loglik -45.15
var- 0.205
a- 75
CED- 1.23
d- 2.48
CES -0.05
CEDL 0.5906
CEDU 2.376
b: -0.03113
conv : 1
scaling factor on x : 1
dtype : 10
selected : all
removed: none
    
```



```

version: 62.6
loglik -45.12
var- 0.205
a- 74.5
CED- 1.48
d- 3.58
CES -0.05
CEDL 0.9195
CEDU 2.382
b: 3.378
conv : 1
scaling factor on x : 1
dtype : 10
selected : all
removed: none
    
```

Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	6	-45.11	102.22
m1-	1	2	-80.52	165.04
m2-	1	3	-51.49	108.98
m3-	1	4	-45.15	98.3
m5-	1	5	-45.11	100.22

AIC: Akaike’s Information Criterion.

Selected model: m3-

The CED (in original units) and the 90 % C.I. is:
 1.2253
 0.5906
 2.3756

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-	1	4	-45.12	98.24
m5-	0	5	-45.11	100.22

AIC: Akaike's Information Criterion.

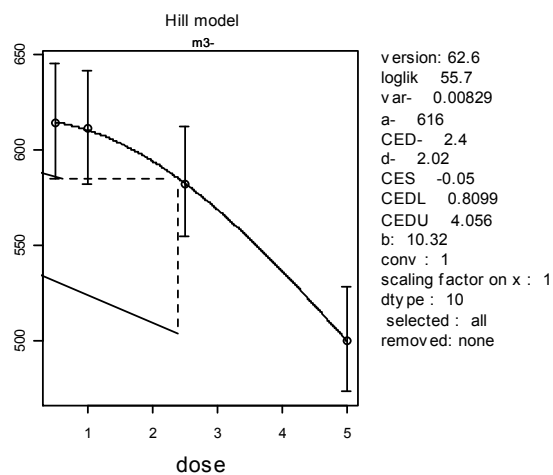
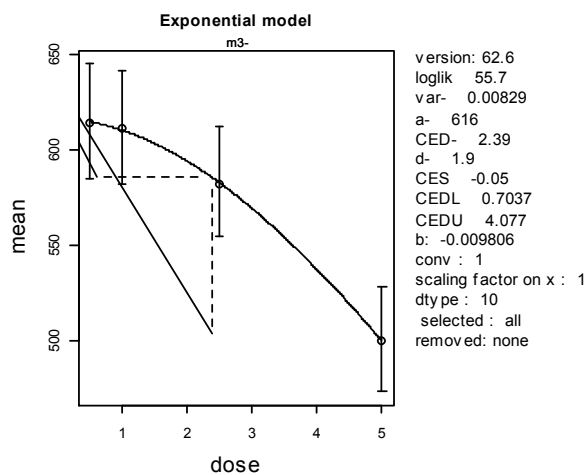
Selected model: m3-

The CED (in original units) and the 90 % C.I. is:
 1.4788
 0.91953
 2.3824

Data, graphic and numerical output for average feed consumption

SprandoAFbis29

5				
0	0	10	0	0
dose	n	mean	sem	sex
0	15	621.3	14.7	m
0.5	15	617.5	17.0	m
1	15	613.6	15.0	m
2.5	15	584.7	13.2	m
5	12	502.1	12.6	m



Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	5	55.71	-101.42
m1-	1	2	39.84	-75.68
m2-	1	3	54.85	-103.7
m3-	1	4	55.7	-103.4
m5-	1	5	55.71	-101.42

AIC: Akaike's Information Criterion.

Selected model: m3-

The CED (in original units) and the 90 % C.I. is:

2.3894
0.70372
4.0772

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-	1	4	55.7	-103.4
m5-	1	5	55.71	-101.42

AIC: Akaike's Information Criterion.

Selected model: m3-

The CED (in original units) and the 90 % C.I. is:

2.405
0.80992
4.0561

Supplementary information COLLINS (Collins et al., 2006)

Details of the BMD Analysis of the data of Collins et al. (2006) reported in Appendix G, Section G.3.2 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

Filename: Collins

Folder: Collins-2006

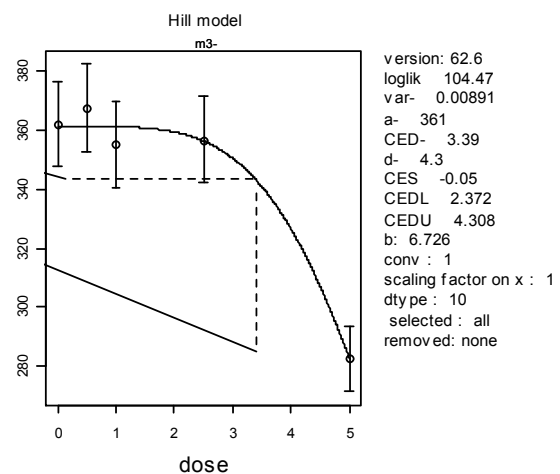
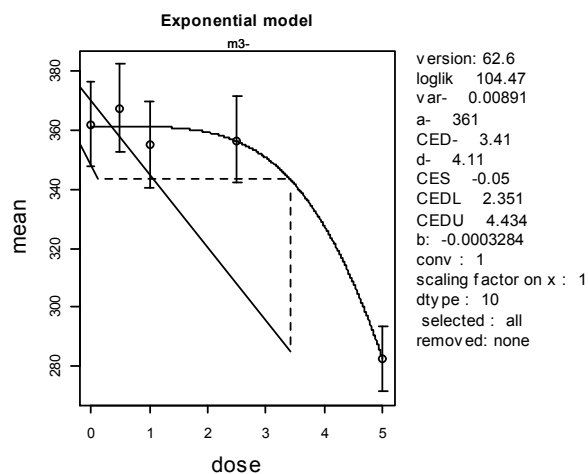
Output from PROAST 62.6 from 24 OCT 2016

Data, graphic and numerical output for body weight

COLLINSBW

5

dose	n	mean	sem	s
0	23	362.22	4.31	s
0.5	22	367.86	4.94	s
1	22	355.86	5.99	s
2.5	21	357.43	5.59	s
5	23	286.22	9.83	s



Response: mean

ANALYSIS WITH EXPONENTIAL MODELS

Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	6	105.2	-198.4
m1-	1	2	63.57	-123.14
m2-	1	3	96.12	-186.24
m3-	1	4	104.47	-200.94
m5-	1	5	104.47	-198.94

AIC: Akaike's Information Criterion.

Selected model: m3-

ATTENTION: jump in log-likelihood

The CED (in original units) and the 90 % C.I. is:

3.4145
2.3508
4.4344

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-	1	4	104.47	-200.94
m5-	1	5	104.47	-198.94

AIC: Akaike's Information Criterion.

Selected model: m3-

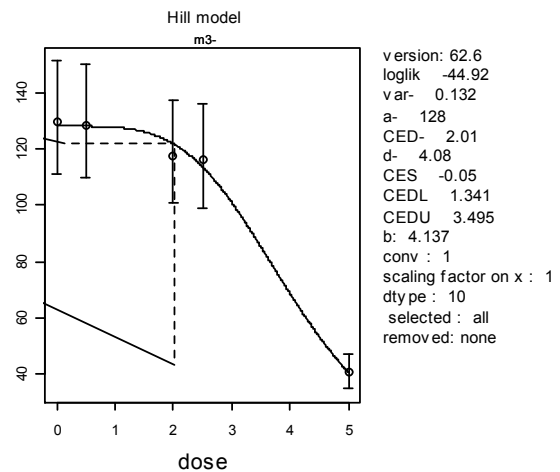
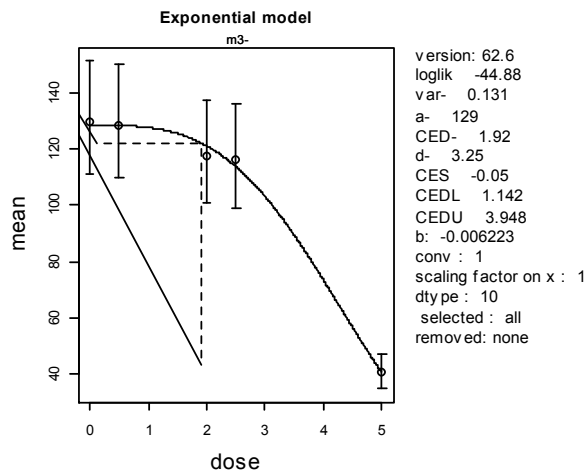
The CED (in original units) and the 90 % C.I. is:

3.393
2.3725
4.3085

Data, graphic and numerical output for body weight gain

COLLINSBWGAIN

```
5
0 0 10 0 0
dose n mean SEM s
0 23 130.7 3.1 s
0.5 22 130.4 4.9 s
2 22 120.8 6 s
2.5 21 118.1 4.9 s
5 23 52.9 9.3 s
```



Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	6	-44.76	101.52
m1-	1	2	-97.03	198.06
m2-	1	3	-58.7	123.4
m3-	1	4	-44.88	97.76
m5-	1	5	-44.88	99.76

AIC: Akaike's Information Criterion.

Selected model: m3-

ATTENTION: jump in log-likelihood

The CED (in original units) and the 90 % C.I. is:
 1.9152
 1.1416
 3.9485

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-	1	4	-44.92	97.84
m5-	1	5	-44.92	99.84

AIC: Akaike's Information Criterion.
 Selected model: m3-

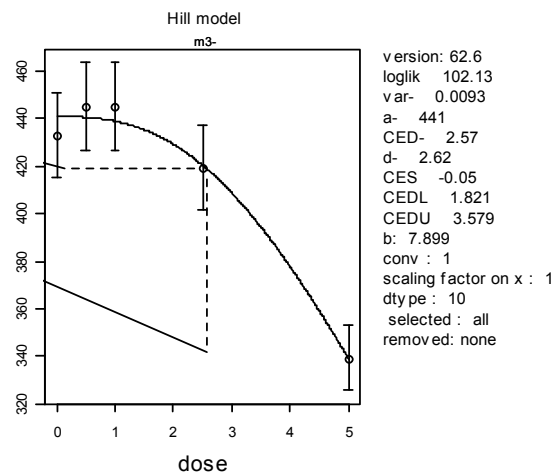
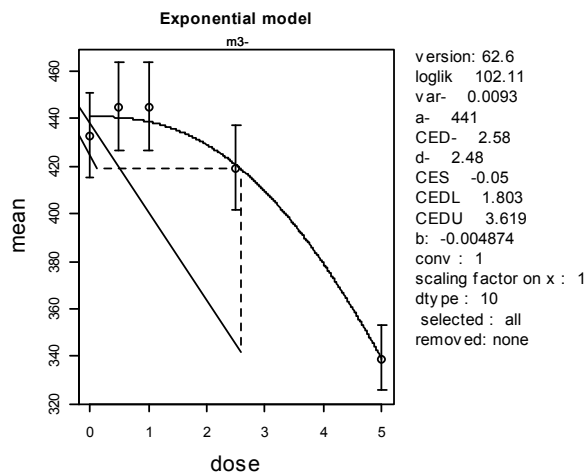
The CED (in original units) and the 90 % C.I. is:
 2.0099
 1.3406
 3.4947

a) Data, graphic and numerical output for feed intake

COLLINSFC

```

5
0 0 10 0 0
dose n mean sem s
0 23 434.3 7.7 s
0.5 22 445.3 5.3 s
1 22 446.1 7.2 s
2.5 21 421.0 9.3 s
5 23 342.6 10.5 s
    
```



Response: mean

ANALYSIS WITH EXPONENTIAL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
full	1	6	102.89	-193.78
m1-	1	2	59.84	-115.68
m2-	1	3	95.25	-184.5
m3-	1	4	102.11	-196.22
m5-	1	5	102.28	-194.56

AIC: Akaike's Information Criterion.

Selected model: m3-

The CED (in original units) and the 90 % C.I. is:

2.5844

1.803

3.6194

Response: mean

ANALYSIS WITH HILL MODELS				
Model	Converged	Number of parameters	Log-likelihood	AIC
m3-	1	4	102.13	-196.26
m5-	1	5	102.28	-194.56

AIC: Akaike's Information Criterion.

Selected model: m3-

The CED (in original units) and the 90 % C.I. is:

2.5722

1.8213

3.5788

Supplementary information YOUNGBMC (Young et al., 1983)

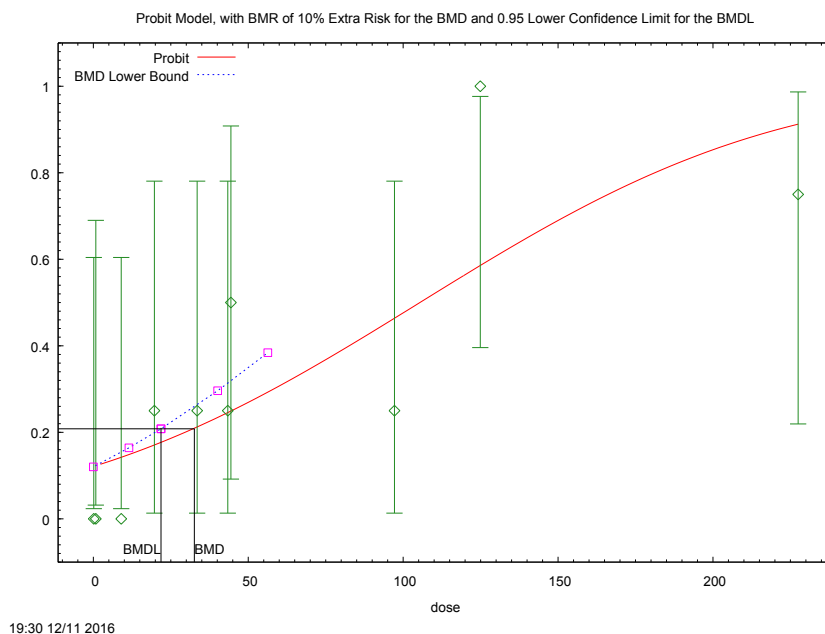
Details of the BMD Analysis of concentration-response data for vomiting in pigs of Young et al. (1983) trials 2 and 3 reported in Appendix G, Section G.4.1.1 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

Filename: YoungBMC Folder: PigsVomiting

Output from BMDS from 11 DEC 2016 with the full information of the fit of all models of BMDS where a model fit was obtained in a reasonable duration of computing time.

The output is ordered in the sequence used in the Table G11 of Appendix G which is based on an order of increasing mathematical complexity. The two less complex multistage models are nested under the multistage model which was used with the default assumption of the degree of the polynomial of BMDS.

Occasionally, it was not possible to obtain the results of the Gamma model with the notebook used for the calculation HP8440p or Lenovo 3508.



```

=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_DONYoungnew_Opt. (d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_DONYoungnew_Opt.plt
Sun Dec 11 19:30:08 2016
=====

```

BMDS_Model_Run

~~~~~

The form of the probability function is:

$$P[\text{response}] = \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Dose}),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect  
 Independent variable = concentration  
 Slope parameter is not restricted

Total number of observations = 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 (and Specified) Parameter Values

background = 0 Specified  
 intercept = -1.18699  
 slope = 0.0110622

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.72 |
| slope     | -0.72     | 1     |

Parameter Estimates

| Interval | Variable  | Estimate  | Std. Err.  | 95.0% Wald Confidence |       |
|----------|-----------|-----------|------------|-----------------------|-------|
|          |           |           |            | Lower Conf. Limit     | Upper |
| 0.523277 | intercept | -1.17418  | 0.332101   | -1.82509              | -     |
| 0.018181 | slope     | 0.0110833 | 0.00362131 | 0.00398569            |       |

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -19.2082        | 2         | 10.3778  | 8         | 0.2395  |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |
| AIC:          | 42.4163         |           |          |           |         |

Goodness of Fit

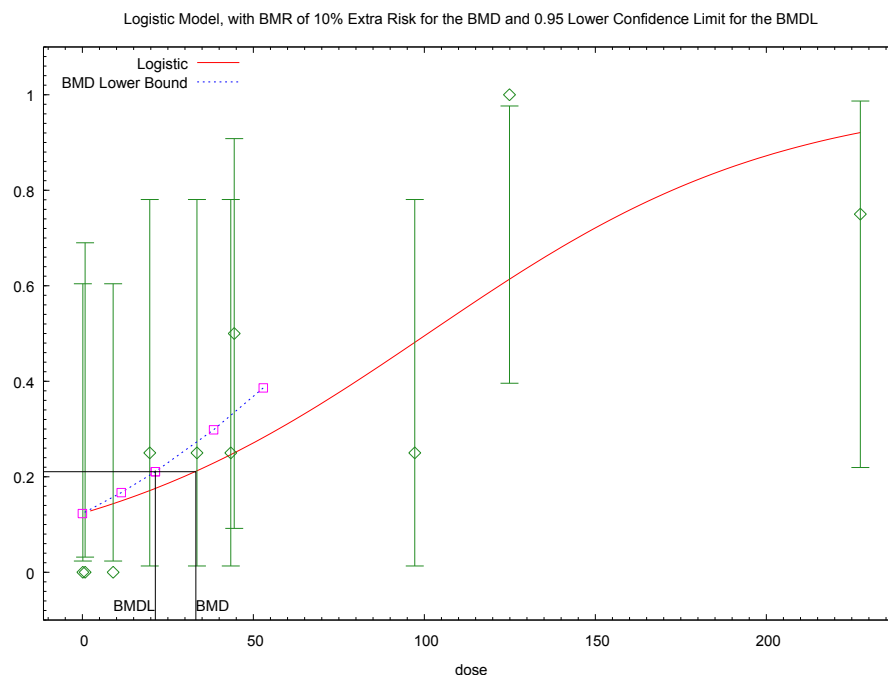
| Dose    | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|---------|------------|----------|----------|-------|-----------------|
| 0.1400  | 0.1205     | 0.482    | 0.000    | 4.000 | -0.740          |
| 9.0000  | 0.1413     | 0.565    | 0.000    | 4.000 | -0.811          |
| 19.7000 | 0.1696     | 0.678    | 1.000    | 4.000 | 0.429           |
| 33.5000 | 0.2110     | 0.844    | 1.000    | 4.000 | 0.191           |
| 43.4000 | 0.2441     | 0.976    | 1.000    | 4.000 | 0.027           |
| 44.4000 | 0.2476     | 0.990    | 2.000    | 4.000 | 1.170           |
| 97.2000 | 0.4614     | 1.846    | 1.000    | 4.000 | -0.848          |

|          |        |       |       |       |        |
|----------|--------|-------|-------|-------|--------|
| 124.9000 | 0.5832 | 2.333 | 4.000 | 4.000 | 1.691  |
| 227.5000 | 0.9111 | 3.644 | 3.000 | 4.000 | -1.132 |
| 0.7900   | 0.1219 | 0.366 | 0.000 | 3.000 | -0.645 |

Chi<sup>2</sup> = 8.07      d.f. = 8      P-value = 0.4267

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 32.5992  
 BMDL = 21.8348



19:33 12/11 2016

```

=====
Logistic Model. (Version: 2.14; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_DONYoungnew_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_DONYoungnew_Opt.plt
Sun Dec 11 19:33:56 2016
=====
    
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = 1/[1+\text{EXP}(-\text{intercept}-\text{slope}*\text{dose})]$$

Dependent variable = Effect  
 Independent variable = concentration  
 Slope parameter is not restricted

Total number of observations = 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  
 background = 0 Specified  
 intercept = -1.55631  
 slope = 0.0147838

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.74 |
| slope     | -0.74     | 1     |

Parameter Estimates

| Interval  | Variable  | Estimate  | Std. Err.  | 95.0% Wald Confidence |       |
|-----------|-----------|-----------|------------|-----------------------|-------|
|           |           |           |            | Lower Conf. Limit     | Upper |
| 0.787695  | intercept | -1.96486  | 0.600604   | -3.14202              | -     |
| 0.0335094 | slope     | 0.0194038 | 0.00719684 | 0.0052983             |       |

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -19.1784        | 2         | 10.3182  | 8         | 0.2434  |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |
| AIC:          | 42.3568         |           |          |           |         |

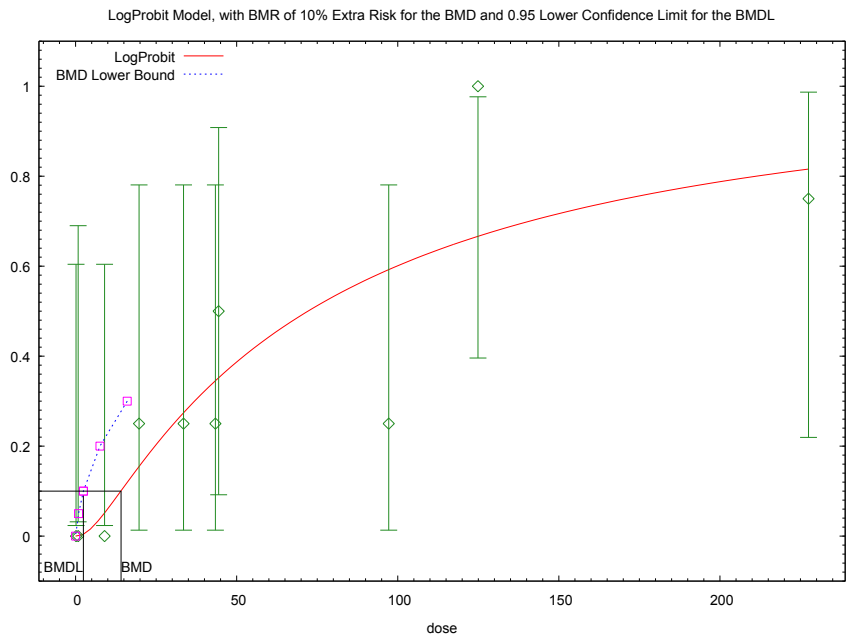
Goodness of Fit

| Dose     | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|----------|------------|----------|----------|-------|-----------------|
| 0.1400   | 0.1232     | 0.493    | 0.000    | 4.000 | -0.750          |
| 9.0000   | 0.1430     | 0.572    | 0.000    | 4.000 | -0.817          |
| 19.7000  | 0.1704     | 0.682    | 1.000    | 4.000 | 0.423           |
| 33.5000  | 0.2117     | 0.847    | 1.000    | 4.000 | 0.188           |
| 43.4000  | 0.2455     | 0.982    | 1.000    | 4.000 | 0.021           |
| 44.4000  | 0.2491     | 0.996    | 2.000    | 4.000 | 1.160           |
| 97.2000  | 0.4803     | 1.921    | 1.000    | 4.000 | -0.922          |
| 124.9000 | 0.6127     | 2.451    | 4.000    | 4.000 | 1.590           |
| 227.5000 | 0.9205     | 3.682    | 3.000    | 4.000 | -1.261          |
| 0.7900   | 0.1246     | 0.374    | 0.000    | 3.000 | -0.653          |

Chi^2 = 8.19      d.f. = 8      P-value = 0.4155

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 33.1808  
 BMDL = 21.3329



19:35 12/11 2016

```
=====  

    Probit Model. (Version: 3.3; Date: 2/28/2013)  

    Input Data File:  

    C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_DONYoungnew_Opt.(d)  

    Gnuplot Plotting File:  

    C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_DONYoungnew_Opt.plt  

    Sun Dec 11 19:35:38 2016  

    =====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{Background} + (1 - \text{Background}) * \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Log}(\text{Dose})),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect  
 Independent variable = concentration  
 Slope parameter is not restricted

Total number of observations = 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



User has chosen the log transformed model

Default Initial (and Specified) Parameter Values

background = 0  
 intercept = -1.20344  
 slope = 0.261415

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.98 |
| slope     | -0.98     | 1     |

Parameter Estimates

|             |            | 95.0% Wald Confidence |           |                   |       |
|-------------|------------|-----------------------|-----------|-------------------|-------|
| Interval    | Variable   | Estimate              | Std. Err. | Lower Conf. Limit | Upper |
| Conf. Limit | background | 0                     | NA        |                   |       |
| 0.996006    | intercept  | -3.35488              | 1.20353   | -5.71375          | -     |
| 1.35504     | slope      | 0.783306              | 0.291705  | 0.211574          |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -17.2699        | 2         | 6.50131  | 8         | 0.5913  |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |

AIC: 38.5399

Goodness of Fit

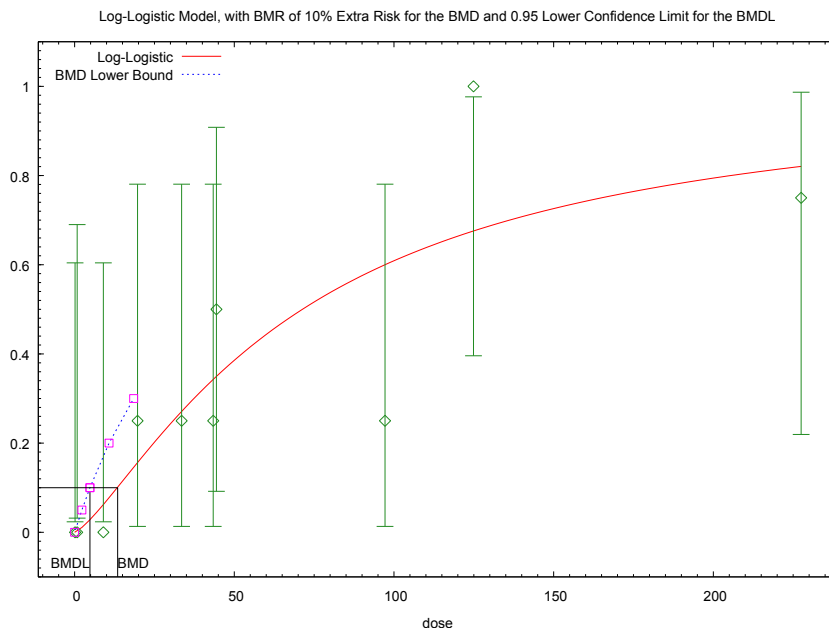
| Dose     | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|----------|------------|----------|----------|-------|-----------------|
| 0.1400   | 0.0000     | 0.000    | 0.000    | 4.000 | -0.001          |
| 9.0000   | 0.0512     | 0.205    | 0.000    | 4.000 | -0.464          |
| 19.7000  | 0.1538     | 0.615    | 1.000    | 4.000 | 0.533           |
| 33.5000  | 0.2728     | 1.091    | 1.000    | 4.000 | -0.103          |
| 43.4000  | 0.3440     | 1.376    | 1.000    | 4.000 | -0.396          |
| 44.4000  | 0.3506     | 1.403    | 2.000    | 4.000 | 0.626           |
| 97.2000  | 0.5910     | 2.364    | 1.000    | 4.000 | -1.387          |
| 124.9000 | 0.6651     | 2.661    | 4.000    | 4.000 | 1.419           |
| 227.5000 | 0.8149     | 3.260    | 3.000    | 4.000 | -0.334          |
| 0.7900   | 0.0002     | 0.001    | 0.000    | 3.000 | -0.025          |

Chi<sup>2</sup> = 5.11      d.f. = 8      P-value = 0.7458

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 14.1101  
 BMDL = 2.41926

Log-Logistic restricted



19:37 12/11 2016

```
=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_DONYoungnew_Opt.(d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_DONYoungnew_Opt.plt
      Sun Dec 11 19:37:27 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect  
 Independent variable = concentration  
 Slope parameter is restricted as slope >= 1

Total number of observations = 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

User has chosen the log transformed model

Default Initial Parameter Values

```
background = 0
intercept = -4.65793
slope = 1
```

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.98 |
| slope     | -0.98     | 1     |

Parameter Estimates

| Interval    | Variable   | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|----------|-----------|-----------------------|-------|
|             |            |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | background | 0        | NA        |                       |       |
| 1.46407     | intercept  | -5.59213 | 2.1062    | -9.7202               | -     |
| 2.30082     | slope      | 1.30533  | 0.507913  | 0.309842              |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -17.316         | 2         | 6.5934   | 8         | 0.5811  |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |

AIC: 38.632

Goodness of Fit

| Dose     | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|----------|------------|----------|----------|-------|-----------------|
| 0.1400   | 0.0003     | 0.001    | 0.000    | 4.000 | -0.034          |
| 9.0000   | 0.0616     | 0.246    | 0.000    | 4.000 | -0.512          |
| 19.7000  | 0.1543     | 0.617    | 1.000    | 4.000 | 0.530           |
| 33.5000  | 0.2673     | 1.069    | 1.000    | 4.000 | -0.078          |
| 43.4000  | 0.3384     | 1.354    | 1.000    | 4.000 | -0.374          |
| 44.4000  | 0.3451     | 1.380    | 2.000    | 4.000 | 0.652           |
| 97.2000  | 0.5944     | 2.377    | 1.000    | 4.000 | -1.403          |
| 124.9000 | 0.6703     | 2.681    | 4.000    | 4.000 | 1.403           |

|          |        |       |       |       |        |
|----------|--------|-------|-------|-------|--------|
| 227.5000 | 0.8164 | 3.266 | 3.000 | 4.000 | -0.343 |
| 0.7900   | 0.0027 | 0.008 | 0.000 | 3.000 | -0.091 |

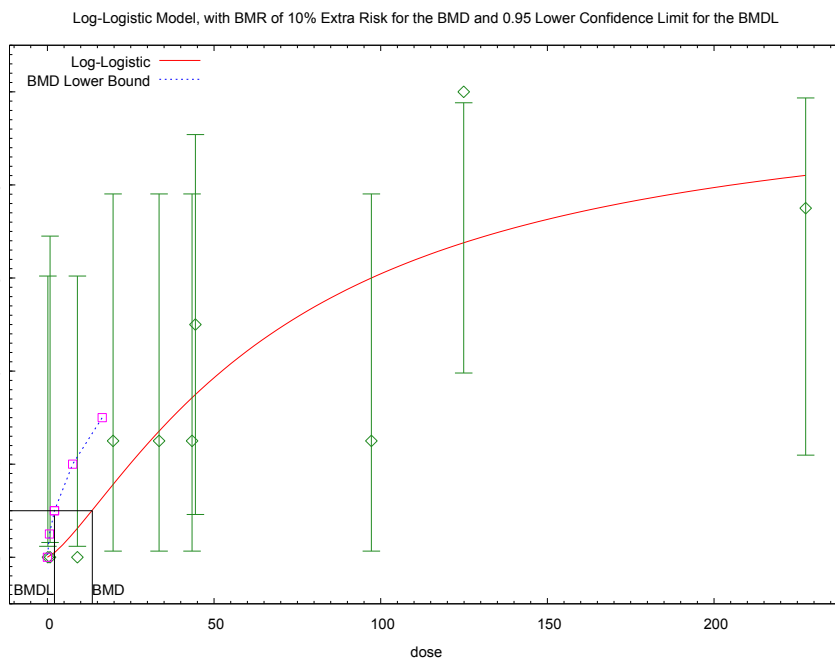
Chi^2 = 5.18      d.f. = 8      P-value = 0.7386

Benchmark Dose Computation

```

Specified effect =          0.1
Risk Type       =          Extra risk
Confidence level =          0.95
                BMD =          13.4745
                BMDL =          4.79127
    
```

loglog unrestricted



19:40 12/11 2016

```

=====
Logistic Model. (Version: 2.14; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_DONYoungnew_Opt. (d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_DONYoungnew_Opt.plt
Sun Dec 11 19:40:26 2016
=====
    
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect  
 Independent variable = concentration

Slope parameter is not restricted

Total number of observations = 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

User has chosen the log transformed model

Default Initial Parameter Values

background = 0  
 intercept = -2.02702  
 slope = 0.444252

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.98 |
| slope     | -0.98     | 1     |

Parameter Estimates

| Interval    | Variable   | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|----------|-----------|-----------------------|-------|
|             |            |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | background | 0        | NA        |                       |       |
| 1.46403     | intercept  | -5.59213 | 2.10621   | -9.72024              | -     |
| 2.30083     | slope      | 1.30533  | 0.507918  | 0.309832              |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -17.316         | 2         | 6.5934   | 8         | 0.5811  |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |
| AIC:          | 38.632          |           |          |           |         |

Goodness of Fit

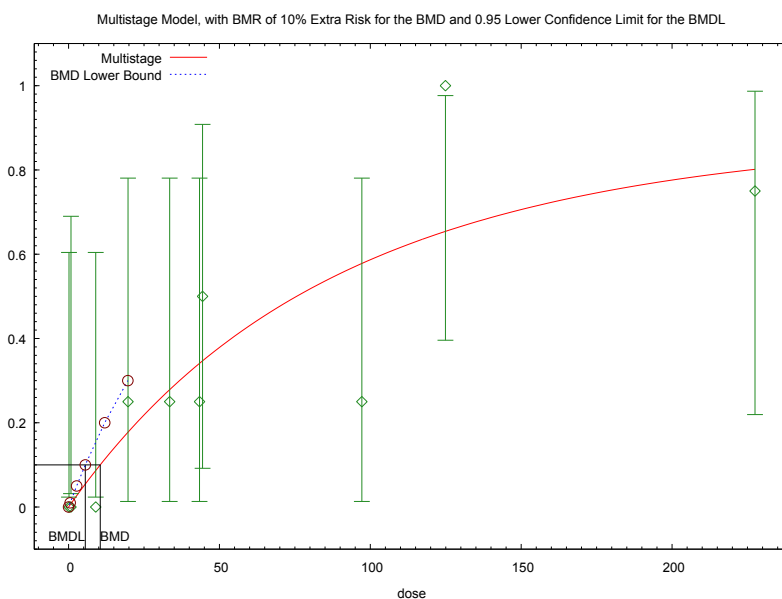
| Dose   | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|--------|------------|----------|----------|-------|-----------------|
| 0.1400 | 0.0003     | 0.001    | 0.000    | 4.000 | -0.034          |
| 9.0000 | 0.0616     | 0.246    | 0.000    | 4.000 | -0.512          |

|          |        |       |       |       |        |
|----------|--------|-------|-------|-------|--------|
| 19.7000  | 0.1543 | 0.617 | 1.000 | 4.000 | 0.530  |
| 33.5000  | 0.2673 | 1.069 | 1.000 | 4.000 | -0.078 |
| 43.4000  | 0.3384 | 1.354 | 1.000 | 4.000 | -0.374 |
| 44.4000  | 0.3451 | 1.380 | 2.000 | 4.000 | 0.652  |
| 97.2000  | 0.5944 | 2.377 | 1.000 | 4.000 | -1.403 |
| 124.9000 | 0.6703 | 2.681 | 4.000 | 4.000 | 1.403  |
| 227.5000 | 0.8164 | 3.266 | 3.000 | 4.000 | -0.343 |
| 0.7900   | 0.0027 | 0.008 | 0.000 | 3.000 | -0.091 |

Chi<sup>2</sup> = 5.18      d.f. = 8      P-value = 0.7386

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 13.4745  
 BMDL = 2.11548



19:38 12/11 2016

```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_DONYoungnew_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_DONYoungnew_Opt.plt
Sun Dec 11 19:38:57 2016
=====
    
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{beta1} * \text{dose}^1 - \text{beta2} * \text{dose}^2)]$$

The parameter betas are not restricted

Dependent variable = Effect  
 Independent variable = concentration

Total number of observations = 10  
 Total number of records with missing values = 0  
 Total number of parameters in model = 3  
 Total number of specified parameters = 0  
 Degree of polynomial = 2

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

Background = 0  
 Beta(1) = 7.96559e+017  
 Beta(2) = -3.05595e+015

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|         | Beta(1) | Beta(2) |
|---------|---------|---------|
| Beta(1) | 1       | -0.86   |
| Beta(2) | -0.86   | 1       |

Parameter Estimates

| Interval     | Variable   | Estimate      | Std. Err.    | 95.0% Wald Confidence |       |
|--------------|------------|---------------|--------------|-----------------------|-------|
|              |            |               |              | Lower Conf. Limit     | Upper |
| Conf. Limit  | Background | 0             | NA           |                       |       |
| 0.0195564    | Beta(1)    | 0.0101776     | 0.00478519   | 0.000798821           |       |
| 5.07281e-005 | Beta(2)    | -1.35925e-005 | 3.28172e-005 | -7.79131e-005         |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -17.3763        | 2         | 6.71407  | 8         | 0.5678  |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |
| AIC:          | 38.7526         |           |          |           |         |

Goodness of Fit

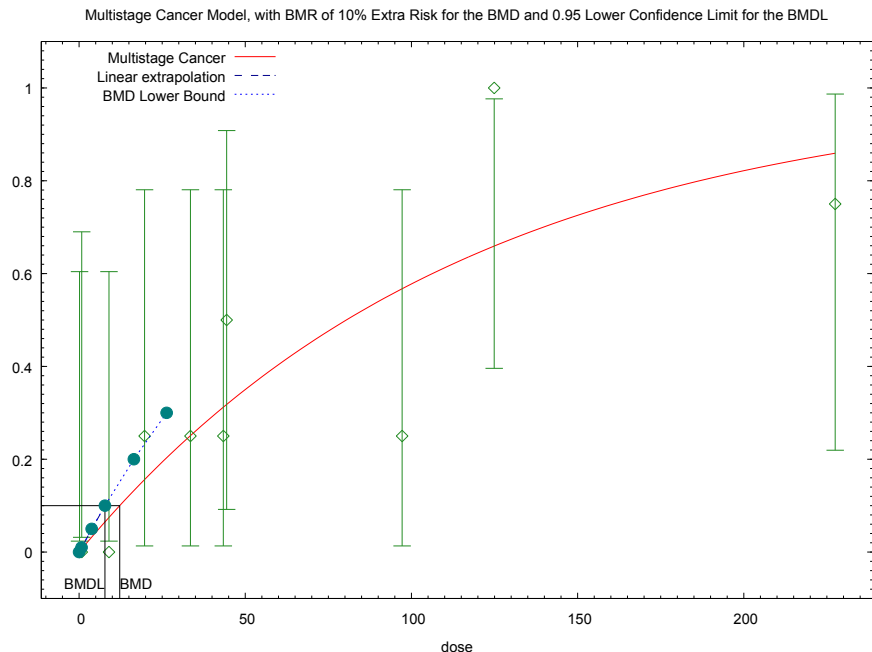
| Dose     | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|----------|------------|----------|----------|-------|-----------------|
| 0.1400   | 0.0014     | 0.006    | 0.000    | 4.000 | -0.076          |
| 9.0000   | 0.0865     | 0.346    | 0.000    | 4.000 | -0.616          |
| 19.7000  | 0.1773     | 0.709    | 1.000    | 4.000 | 0.380           |
| 33.5000  | 0.2780     | 1.112    | 1.000    | 4.000 | -0.125          |
| 43.4000  | 0.3404     | 1.362    | 1.000    | 4.000 | -0.382          |
| 44.4000  | 0.3463     | 1.385    | 2.000    | 4.000 | 0.646           |
| 97.2000  | 0.5772     | 2.309    | 1.000    | 4.000 | -1.325          |
| 124.9000 | 0.6532     | 2.613    | 4.000    | 4.000 | 1.457           |
| 227.5000 | 0.8005     | 3.202    | 3.000    | 4.000 | -0.253          |
| 0.7900   | 0.0080     | 0.024    | 0.000    | 3.000 | -0.156          |

Chi<sup>2</sup> = 5.07      d.f. = 8      P-value = 0.7496

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 10.4994  
 BMDL = 5.53894  
 BMDU = 27.9637

Taken together, (5.53894, 27.9637) is a 90 % two-sided confidence interval for the BMD



19:42 12/11 2016

```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_DONYoungnew_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_DONYoungnew_Opt.plt
Sun Dec 11 19:42:25 2016
    
```



=====

BMDS\_Model\_Run

~~~~~

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1 - \text{EXP}(-\text{beta1} * \text{dose}^{\text{beta2}})]$$

The parameter betas are restricted to be positive

Dependent variable = Effect
Independent variable = concentration

Total number of observations = 10
Total number of records with missing values = 0
Total number of parameters in model = 3
Total number of specified parameters = 0
Degree of polynomial = 2

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

Background = 1
Beta(1) = 1.404e+017
Beta(2) = 0

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Beta(2) have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

Beta(1)
Beta(1) 1

Parameter Estimates

Interval		95.0% Wald Confidence		
Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper Conf. Limit
Background	0	NA		
Beta(1)	0.00862046	0.00251679	0.00368764	
Beta(2)	0	NA		

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-14.0193	10			
Fitted model	-17.4576	1	6.87667	9	0.65
Reduced model	-24.8241	1	21.6095	9	0.0102

AIC: 36.9153

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.1400	0.0012	0.005	0.000	4.000	-0.070
9.0000	0.0747	0.299	0.000	4.000	-0.568
19.7000	0.1562	0.625	1.000	4.000	0.517
33.5000	0.2508	1.003	1.000	4.000	-0.004
43.4000	0.3121	1.248	1.000	4.000	-0.268
44.4000	0.3180	1.272	2.000	4.000	0.782
97.2000	0.5674	2.270	1.000	4.000	-1.281
124.9000	0.6593	2.637	4.000	4.000	1.438
227.5000	0.8593	3.437	3.000	4.000	-0.629
0.7900	0.0068	0.020	0.000	3.000	-0.143

Chi² = 5.40 d.f. = 9 P-value = 0.7980

Benchmark Dose Computation

Specified effect = 0.1

Risk Type = Extra risk

Confidence level = 0.95

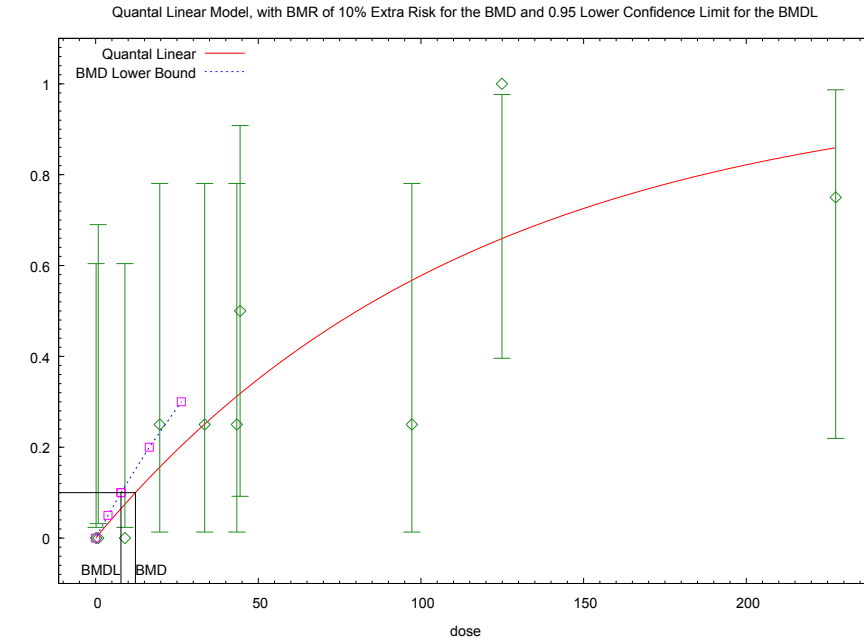
BMD = 12.2221

BMDL = 7.782

BMDU = 29.3638

Taken together, (7.782 , 29.3638) is a 90 % two-sided confidence interval for the BMD

Cancer Slope Factor = 0.0128502



```

=====
      Quantal Linear Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_DONYoungnew_Opt.(d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_DONYoungnew_Opt.plt
      Sun Dec 11 19:44:37 2016
=====

```

BMDS_Model_Run

~~~~~

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) * [1 - \text{EXP}(-\text{slope} * \text{dose})]$$

Dependent variable = Effect  
 Independent variable = concentration

Total number of observations = 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 (and Specified) Parameter Values

Background = 0.0833333  
 Slope = 0.0044466  
 Power = 1 Specified

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background -Power  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

Slope  
Slope 1

Parameter Estimates

95.0% Wald Confidence

| Interval    | Variable   | Estimate   | Std. Err.  | Lower Conf. Limit | Upper |
|-------------|------------|------------|------------|-------------------|-------|
| Conf. Limit | Background | 0          | NA         |                   |       |
| 0.0135533   | Slope      | 0.00862046 | 0.00251679 | 0.00368764        |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -17.4576        | 1         | 6.87667  | 9         | 0.65    |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |

AIC: 36.9153

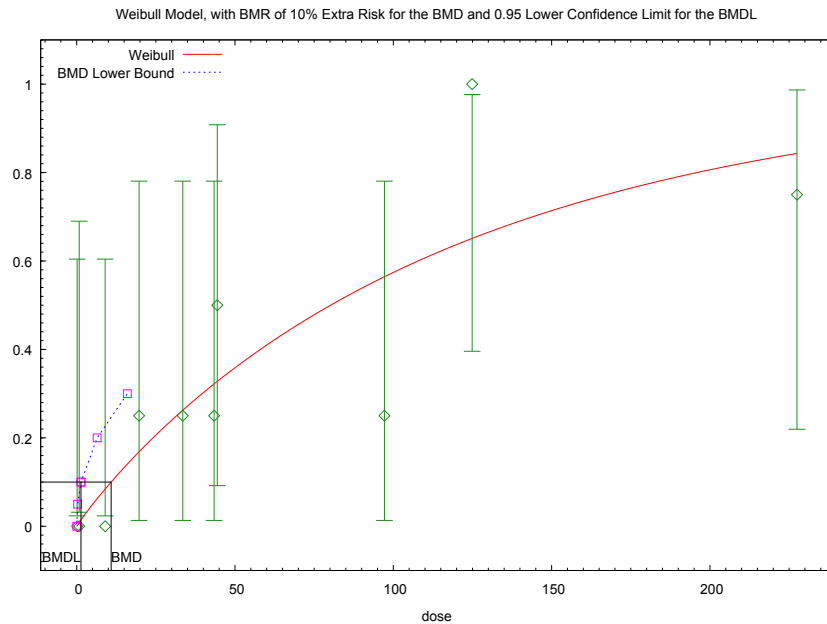
Goodness of Fit

| Dose     | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|----------|------------|----------|----------|-------|-----------------|
| 0.1400   | 0.0012     | 0.005    | 0.000    | 4.000 | -0.070          |
| 9.0000   | 0.0747     | 0.299    | 0.000    | 4.000 | -0.568          |
| 19.7000  | 0.1562     | 0.625    | 1.000    | 4.000 | 0.517           |
| 33.5000  | 0.2508     | 1.003    | 1.000    | 4.000 | -0.004          |
| 43.4000  | 0.3121     | 1.248    | 1.000    | 4.000 | -0.268          |
| 44.4000  | 0.3180     | 1.272    | 2.000    | 4.000 | 0.782           |
| 97.2000  | 0.5674     | 2.270    | 1.000    | 4.000 | -1.281          |
| 124.9000 | 0.6593     | 2.637    | 4.000    | 4.000 | 1.438           |
| 227.5000 | 0.8593     | 3.437    | 3.000    | 4.000 | -0.629          |
| 0.7900   | 0.0068     | 0.020    | 0.000    | 3.000 | -0.143          |

Chi<sup>2</sup> = 5.40      d.f. = 9      P-value = 0.7980

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 12.2221  
 BMDL = 7.78199



19:46 12/11 2016

```
=====
Weibull Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_DONYoungnew_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_DONYoungnew_Opt.plt
Sun Dec 11 19:46:05 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose}^{\text{power}})]$$

Dependent variable = Effect  
 Independent variable = concentration  
 Power parameter is not restricted

Total number of observations = 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 (and Specified) Parameter Values

Background = 0.0833333  
 Slope = 0.0415303  
 Power = 0.588314

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

Slope                  Power

|       |       |       |
|-------|-------|-------|
| Slope | 1     | -0.98 |
| Power | -0.98 | 1     |

Parameter Estimates

|             |            |           |                       |                   |       |
|-------------|------------|-----------|-----------------------|-------------------|-------|
|             |            |           | 95.0% Wald Confidence |                   |       |
| Interval    | Variable   | Estimate  | Std. Err.             | Lower Conf. Limit | Upper |
| Conf. Limit | Background | 0         | NA                    |                   |       |
| 0.0432303   | Slope      | 0.0111071 | 0.0163896             | -0.021016         |       |
| 1.58765     | Power      | 0.942577  | 0.329127              | 0.297501          |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

|               |                 |           |          |           |         |
|---------------|-----------------|-----------|----------|-----------|---------|
| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
| Full model    | -14.0193        | 10        |          |           |         |
| Fitted model  | -17.4427        | 2         | 6.84678  | 8         | 0.5533  |
| Reduced model | -24.8241        | 1         | 21.6095  | 9         | 0.0102  |
| AIC:          | 38.8854         |           |          |           |         |

Goodness of Fit

| Dose     | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|----------|------------|----------|----------|-------|-----------------|
| 0.1400   | 0.0017     | 0.007    | 0.000    | 4.000 | -0.083          |
| 9.0000   | 0.0843     | 0.337    | 0.000    | 4.000 | -0.607          |
| 19.7000  | 0.1684     | 0.674    | 1.000    | 4.000 | 0.436           |
| 33.5000  | 0.2622     | 1.049    | 1.000    | 4.000 | -0.056          |
| 43.4000  | 0.3217     | 1.287    | 1.000    | 4.000 | -0.307          |
| 44.4000  | 0.3274     | 1.310    | 2.000    | 4.000 | 0.736           |
| 97.2000  | 0.5640     | 2.256    | 1.000    | 4.000 | -1.266          |
| 124.9000 | 0.6506     | 2.602    | 4.000    | 4.000 | 1.466           |
| 227.5000 | 0.8428     | 3.371    | 3.000    | 4.000 | -0.510          |
| 0.7900   | 0.0089     | 0.027    | 0.000    | 3.000 | -0.164          |

Chi^2 = 5.24      d.f. = 8      P-value = 0.7313

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 10.8793  
 BMDL = 1.35813

## Supplementary information WILLIAMS BMC (Williams et al., 1988)

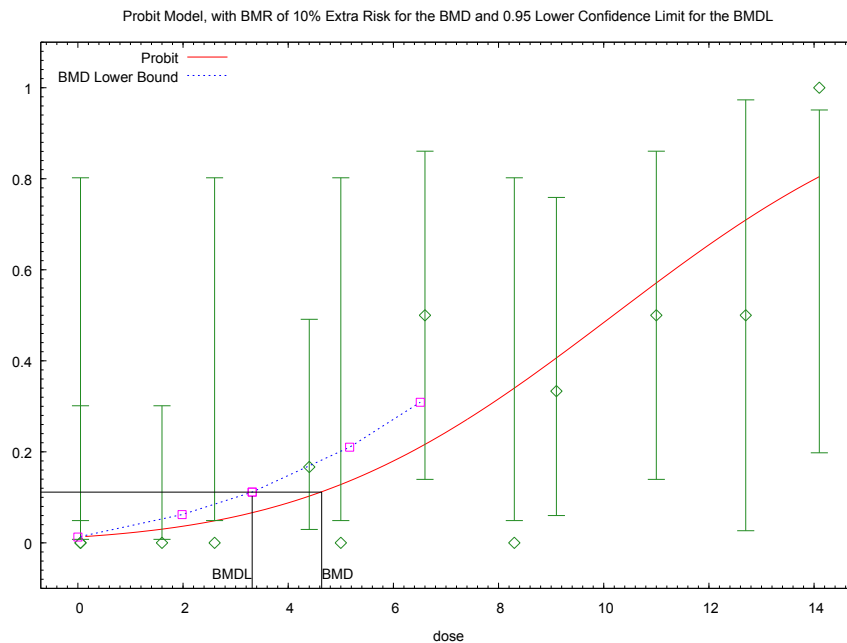
Details of the BMD Analysis of concentration-response data for vomiting in pigs of Williams et al. (1986) trials 1 and 2 reported in Appendix G, Section G.4.1.2 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

Filename: WilliamsBMC Folder: PigsVomiting

Output from BMDS from 11 DEC 2016 with the full information of the fit of all models of BMDS where a model fit was obtained in a reasonable duration of computing time.

The output is ordered in the sequence used in Table G12 shown in Appendix G which is based on an order of increasing mathematical complexity. The two less complex multistage models are nested under the multistage model which was used with the default assumption of the degree of the polynomial of BMDS.

Occasionally, it was not possible to obtain the results of the Gamma model with the notebook used for the calculation HP844op or Lenovo 3508.



```

=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_williams_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_williams_Opt.plt
Sun Dec 11 20:18:30 2016
=====

```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Dose}),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 12  
 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 (and Specified) Parameter Values  
 background = 0 Specified  
 intercept = -2.14661  
 slope = 0.209647

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.89 |
| slope     | -0.89     | 1     |

Parameter Estimates

| Interval    | Variable  | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|-----------|----------|-----------|-----------------------|-------|
|             |           |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | intercept | -2.23445 | 0.465752  | -3.1473               | -     |
| 1.32159     | slope     | 0.219064 | 0.0576327 | 0.106106              |       |
| 0.332022    |           |          |           |                       |       |

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value  |
|---------------|-----------------|-----------|----------|-----------|----------|
| Full model    | -18.9299        | 12        |          |           |          |
| Fitted model  | -22.8382        | 2         | 7.81667  | 10        | 0.6467   |
| Reduced model | -32.7474        | 1         | 27.635   | 11        | 0.003681 |
| AIC:          | 49.6764         |           |          |           |          |

Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size   | Scaled Residual |
|--------|------------|----------|----------|--------|-----------------|
| 0.0500 | 0.0131     | 0.157    | 0.000    | 12.000 | -0.399          |
| 1.6000 | 0.0298     | 0.357    | 0.000    | 12.000 | -0.607          |
| 4.4000 | 0.1019     | 1.223    | 2.000    | 12.000 | 0.741           |
| 6.6000 | 0.2152     | 1.291    | 3.000    | 6.000  | 1.698           |
| 9.1000 | 0.4048     | 2.429    | 2.000    | 6.000  | -0.357          |



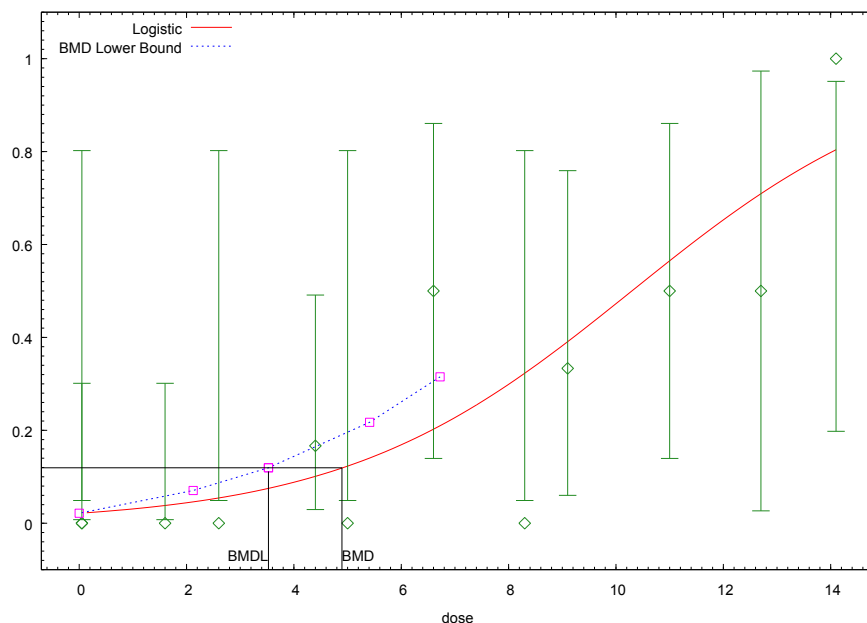
|         |        |       |       |       |        |
|---------|--------|-------|-------|-------|--------|
| 11.0000 | 0.5696 | 3.417 | 3.000 | 6.000 | -0.344 |
| 0.0500  | 0.0131 | 0.026 | 0.000 | 2.000 | -0.163 |
| 2.6000  | 0.0480 | 0.096 | 0.000 | 2.000 | -0.317 |
| 5.0000  | 0.1273 | 0.255 | 0.000 | 2.000 | -0.540 |
| 8.3000  | 0.3386 | 0.677 | 0.000 | 2.000 | -1.012 |
| 12.7000 | 0.7080 | 1.416 | 1.000 | 2.000 | -0.647 |
| 14.1000 | 0.8035 | 1.607 | 2.000 | 2.000 | 0.699  |

Chi<sup>2</sup> = 6.56      d.f. = 10      P-value = 0.7666

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 4.63617  
 BMDL = 3.31477

Logistic Model, with BMR of 10% Extra Risk for the BMD and 0.95 Lower Confidence Limit for the BMDL



20:21 12/11 2016

```

=====
Logistic Model. (Version: 2.14; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_williams_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_williams_Opt.plt
Sun Dec 11 20:21:12 2016
=====
    
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = 1/[1+\text{EXP}(-\text{intercept}-\text{slope}*\text{dose})]$$

Dependent variable = Effect  
 Independent variable = Dose

Slope parameter is not restricted

Total number of observations = 12  
 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  
 background = 0 Specified  
 intercept = -2.6779  
 slope = 0.249614

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.91 |
| slope     | -0.91     | 1     |

Parameter Estimates

| Interval    | Variable  | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|-----------|----------|-----------|-----------------------|-------|
|             |           |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | intercept | -3.81662 | 0.884678  | -5.55056              | -     |
| 2.08269     | slope     | 0.371464 | 0.103439  | 0.168728              |       |
| 0.574201    |           |          |           |                       |       |

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value  |
|---------------|-----------------|-----------|----------|-----------|----------|
| Full model    | -18.9299        | 12        |          |           |          |
| Fitted model  | -23.1391        | 2         | 8.41841  | 10        | 0.588    |
| Reduced model | -32.7474        | 1         | 27.635   | 11        | 0.003681 |
| AIC:          | 50.2782         |           |          |           |          |

Goodness of Fit

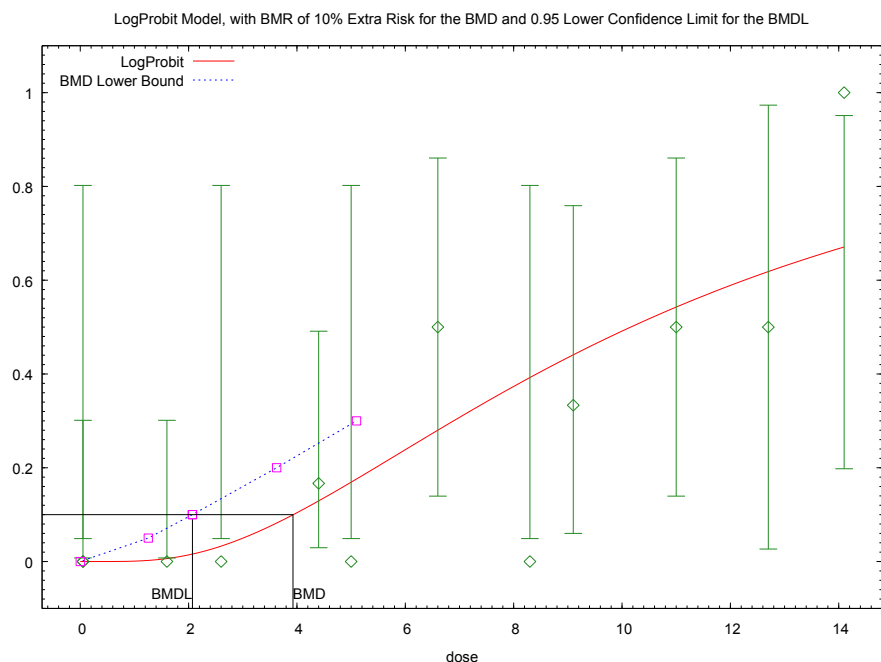
| Dose    | Est._Prob. | Expected | Observed | Size   | Scaled Residual |
|---------|------------|----------|----------|--------|-----------------|
| 0.0500  | 0.0219     | 0.263    | 0.000    | 12.000 | -0.519          |
| 1.6000  | 0.0383     | 0.460    | 0.000    | 12.000 | -0.692          |
| 4.4000  | 0.1014     | 1.216    | 2.000    | 12.000 | 0.750           |
| 6.6000  | 0.2034     | 1.221    | 3.000    | 6.000  | 1.805           |
| 9.1000  | 0.3926     | 2.356    | 2.000    | 6.000  | -0.297          |
| 11.0000 | 0.5670     | 3.402    | 3.000    | 6.000  | -0.331          |
| 0.0500  | 0.0219     | 0.044    | 0.000    | 2.000  | -0.212          |
| 2.6000  | 0.0546     | 0.109    | 0.000    | 2.000  | -0.340          |
| 5.0000  | 0.1235     | 0.247    | 0.000    | 2.000  | -0.531          |
| 8.3000  | 0.3244     | 0.649    | 0.000    | 2.000  | -0.980          |

|         |        |       |       |       |        |
|---------|--------|-------|-------|-------|--------|
| 12.7000 | 0.7111 | 1.422 | 1.000 | 2.000 | -0.659 |
| 14.1000 | 0.8055 | 1.611 | 2.000 | 2.000 | 0.695  |

Chi<sup>2</sup> = 7.08      d.f. = 10      P-value = 0.7175

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 4.89486  
 BMDL = 3.52599



20:23 12/11 2016

```

=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_williams_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_williams_Opt.plt
Sun Dec 11 20:23:39 2016
=====
  
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{Background} + (1 - \text{Background}) * \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Log}(\text{Dose})),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 12  
 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

User has chosen the log transformed model

Default Initial (and Specified) Parameter Values  
 background = 0  
 intercept = -1.11261  
 slope = 0.273357

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.97 |
| slope     | -0.97     | 1     |

Parameter Estimates

| Interval    | Variable   | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|----------|-----------|-----------------------|-------|
|             |            |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | background | 0        | NA        |                       |       |
| 1.28891     | intercept  | -3.12961 | 0.93915   | -4.97031              | -     |
| 2.25991     | slope      | 1.3512   | 0.463634  | 0.442499              |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value  |
|---------------|-----------------|-----------|----------|-----------|----------|
| Full model    | -18.9299        | 12        |          |           |          |
| Fitted model  | -22.1758        | 2         | 6.49181  | 10        | 0.7724   |
| Reduced model | -32.7474        | 1         | 27.635   | 11        | 0.003681 |
| AIC:          | 48.3516         |           |          |           |          |

Goodness of Fit

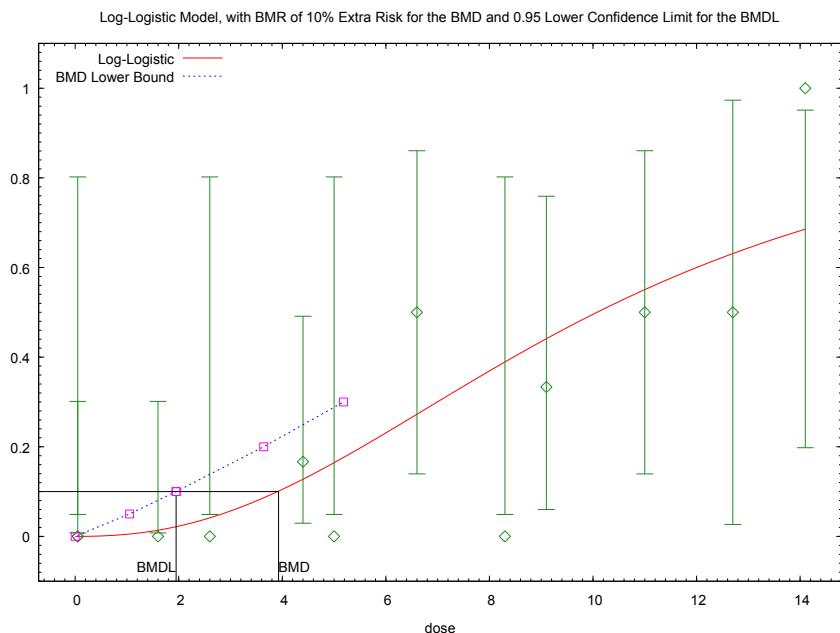
| Dose   | Est._Prob. | Expected | Observed | Size   | Scaled Residual |
|--------|------------|----------|----------|--------|-----------------|
| 0.0500 | 0.0000     | 0.000    | 0.000    | 12.000 | -0.000          |
| 1.6000 | 0.0063     | 0.076    | 0.000    | 12.000 | -0.276          |
| 4.4000 | 0.1297     | 1.557    | 2.000    | 12.000 | 0.381           |
| 6.6000 | 0.2810     | 1.686    | 3.000    | 6.000  | 1.193           |

|         |        |       |       |       |        |
|---------|--------|-------|-------|-------|--------|
| 9.1000  | 0.4420 | 2.652 | 2.000 | 6.000 | -0.536 |
| 11.0000 | 0.5440 | 3.264 | 3.000 | 6.000 | -0.216 |
| 0.0500  | 0.0000 | 0.000 | 0.000 | 2.000 | -0.000 |
| 2.6000  | 0.0330 | 0.066 | 0.000 | 2.000 | -0.261 |
| 5.0000  | 0.1698 | 0.340 | 0.000 | 2.000 | -0.640 |
| 8.3000  | 0.3935 | 0.787 | 0.000 | 2.000 | -1.139 |
| 12.7000 | 0.6197 | 1.239 | 1.000 | 2.000 | -0.349 |
| 14.1000 | 0.6722 | 1.344 | 2.000 | 2.000 | 0.988  |

Chi<sup>2</sup> = 4.85      d.f. = 10      P-value = 0.9009

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 3.92637  
 BMDL = 2.07069



20:25 12/11 2016

```
=====
Logistic Model. (Version: 2.14; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_williams_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_williams_Opt.plt
Sun Dec 11 20:25:16 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect  
 Independent variable = Dose

Slope parameter is not restricted

Total number of observations = 12  
 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

User has chosen the log transformed model

Default Initial Parameter Values

background = 0  
 intercept = -1.59731  
 slope = 0.454382

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.98 |
| slope     | -0.98     | 1     |

Parameter Estimates

| Interval    | Variable   | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|----------|-----------|-----------------------|-------|
|             |            |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | background | 0        | NA        |                       |       |
| 2.03048     | intercept  | -5.36433 | 1.70098   | -8.69819              | -     |
| 3.91732     | slope      | 2.31515  | 0.817445  | 0.712991              |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value  |
|---------------|-----------------|-----------|----------|-----------|----------|
| Full model    | -18.9299        | 12        |          |           |          |
| Fitted model  | -22.3001        | 2         | 6.74039  | 10        | 0.7497   |
| Reduced model | -32.7474        | 1         | 27.635   | 11        | 0.003681 |
| AIC:          | 48.6001         |           |          |           |          |

Goodness of Fit

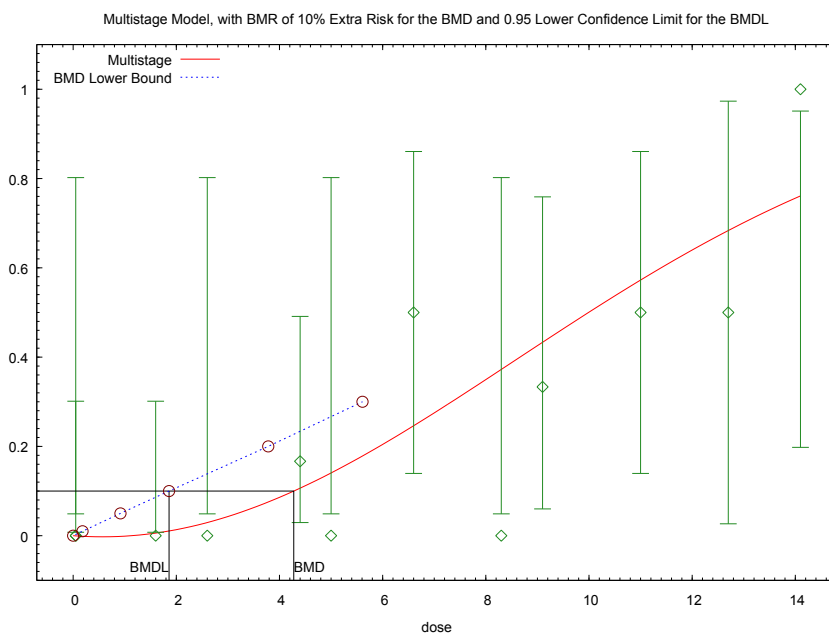
| Dose   | Est._Prob. | Expected | Observed | Size   | Scaled Residual |
|--------|------------|----------|----------|--------|-----------------|
| 0.0500 | 0.0000     | 0.000    | 0.000    | 12.000 | -0.007          |
| 1.6000 | 0.0137     | 0.164    | 0.000    | 12.000 | -0.408          |

|         |        |       |       |        |        |
|---------|--------|-------|-------|--------|--------|
| 4.4000  | 0.1263 | 1.515 | 2.000 | 12.000 | 0.421  |
| 6.6000  | 0.2698 | 1.619 | 3.000 | 6.000  | 1.270  |
| 9.1000  | 0.4374 | 2.624 | 2.000 | 6.000  | -0.514 |
| 11.0000 | 0.5467 | 3.280 | 3.000 | 6.000  | -0.230 |
| 0.0500  | 0.0000 | 0.000 | 0.000 | 2.000  | -0.003 |
| 2.6000  | 0.0410 | 0.082 | 0.000 | 2.000  | -0.292 |
| 5.0000  | 0.1627 | 0.325 | 0.000 | 2.000  | -0.623 |
| 8.3000  | 0.3858 | 0.772 | 0.000 | 2.000  | -1.121 |
| 12.7000 | 0.6271 | 1.254 | 1.000 | 2.000  | -0.372 |
| 14.1000 | 0.6818 | 1.364 | 2.000 | 2.000  | 0.966  |

Chi<sup>2</sup> = 5.08      d.f. = 10      P-value = 0.8860

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 3.92745  
 BMDL = 1.94923



20:27 12/11 2016

```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_williams_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_williams_Opt.plt
Sun Dec 11 20:27:50 2016
=====
    
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{beta1} * \text{dose}^{1-\text{beta2}} * \text{dose}^2)]$$

The parameter betas are not restricted

Dependent variable = Effect  
Independent variable = Dose

Total number of observations = 12  
Total number of records with missing values = 0  
Total number of parameters in model = 3  
Total number of specified parameters = 0  
Degree of polynomial = 2

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

Background = 1  
Beta(1) = -7.78141e+018  
Beta(2) = 8.05238e+017

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|         | Beta(1) | Beta(2) |
|---------|---------|---------|
| Beta(1) | 1       | -0.93   |
| Beta(2) | -0.93   | 1       |

Parameter Estimates

| Interval    | Variable   | Estimate    | Std. Err.  | 95.0% Wald Confidence |       |
|-------------|------------|-------------|------------|-----------------------|-------|
|             |            |             |            | Lower Conf. Limit     | Upper |
| Conf. Limit | Background | 0           | NA         |                       |       |
| 0.0584069   | Beta(1)    | -0.00886195 | 0.0343215  | -0.0761308            |       |
| 0.0176364   | Beta(2)    | 0.00783137  | 0.00500266 | -0.00197366           |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value  |
|---------------|-----------------|-----------|----------|-----------|----------|
| Full model    | -18.9299        | 12        |          |           |          |
| Fitted model  | -22.261         | 2         | 6.66215  | 10        | 0.7569   |
| Reduced model | -32.7474        | 1         | 27.635   | 11        | 0.003681 |
| AIC:          | 48.5219         |           |          |           |          |



Goodness of Fit

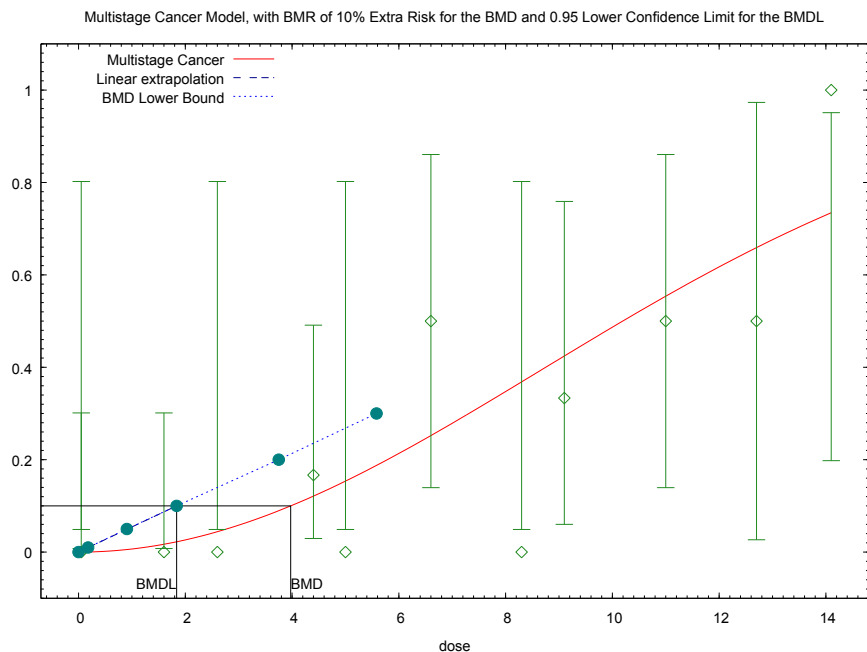
| Dose    | Est._Prob. | Expected | Observed | Size   | Scaled Residual |
|---------|------------|----------|----------|--------|-----------------|
| 0.0500  | -0.0004    | -0.005   | 0.000    | 12.000 | 0.005           |
| 1.6000  | 0.0059     | 0.070    | 0.000    | 12.000 | -0.266          |
| 4.4000  | 0.1065     | 1.278    | 2.000    | 12.000 | 0.675           |
| 6.6000  | 0.2462     | 1.477    | 3.000    | 6.000  | 1.443           |
| 9.1000  | 0.4333     | 2.600    | 2.000    | 6.000  | -0.494          |
| 11.0000 | 0.5726     | 3.436    | 3.000    | 6.000  | -0.360          |
| 0.0500  | -0.0004    | -0.001   | 0.000    | 2.000  | 0.001           |
| 2.6000  | 0.0295     | 0.059    | 0.000    | 2.000  | -0.246          |
| 5.0000  | 0.1406     | 0.281    | 0.000    | 2.000  | -0.572          |
| 8.3000  | 0.3725     | 0.745    | 0.000    | 2.000  | -1.090          |
| 12.7000 | 0.6835     | 1.367    | 1.000    | 2.000  | -0.558          |
| 14.1000 | 0.7612     | 1.522    | 2.000    | 2.000  | 0.792           |

Chi<sup>2</sup> = 5.50      d.f. = 10      P-value = 0.8557

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 4.2771  
 BMDL = 1.85926  
 BMDU = 5.85534

Taken together, (1.85926, 5.85534) is a 90 % two-sided confidence interval for the BMD



20:29 12/11 2016

=====  
 Multistage Model. (Version: 3.4; Date: 05/02/2014)  
 Input Data File:  
 C:/Users/edler/Desktop/BMDS2601\_20150629/BMDS2601/Data/msc\_williams\_Opt.(d)

Gnuplot Plotting File:  
 C:/Users/edler/Desktop/BMDS2601\_20150629/BMDS2601/Data/msc\_williams\_Opt.plt  
 Sun Dec 11 20:29:44 2016

=====

BMDS\_Model\_Run

~~~~~

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1 - \text{EXP}(-\text{beta1} * \text{dose}^{\text{beta2}})]$$

The parameter betas are restricted to be positive

Dependent variable = Effect
 Independent variable = Dose

Total number of observations = 12
 Total number of records with missing values = 0
 Total number of parameters in model = 3
 Total number of specified parameters = 0
 Degree of polynomial = 2

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

Background = 0
 Beta(1) = 0
 Beta(2) = 2.74191e+017

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Beta(1)
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

Beta(2)

Beta(2) 1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
	Beta(1)	0	NA		
	Beta(2)	0.00667323	0.00189447	0.00296013	
		0.0103863			

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-18.9299	12			
Fitted model	-22.2931	1	6.72636	11	0.8208
Reduced model	-32.7474	1	27.635	11	0.003681

AIC: 46.5861

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0500	0.0000	0.000	0.000	12.000	-0.014
1.6000	0.0169	0.203	0.000	12.000	-0.455
4.4000	0.1212	1.454	2.000	12.000	0.483
6.6000	0.2522	1.513	3.000	6.000	1.397
9.1000	0.4246	2.547	2.000	6.000	-0.452
11.0000	0.5540	3.324	3.000	6.000	-0.266
0.0500	0.0000	0.000	0.000	2.000	-0.006
2.6000	0.0441	0.088	0.000	2.000	-0.304
5.0000	0.1537	0.307	0.000	2.000	-0.603
8.3000	0.3685	0.737	0.000	2.000	-1.080
12.7000	0.6592	1.318	1.000	2.000	-0.475
14.1000	0.7346	1.469	2.000	2.000	0.850

Chi² = 5.24 d.f. = 11 P-value = 0.9191

Benchmark Dose Computation

Specified effect = 0.1

Risk Type = Extra risk

Confidence level = 0.95

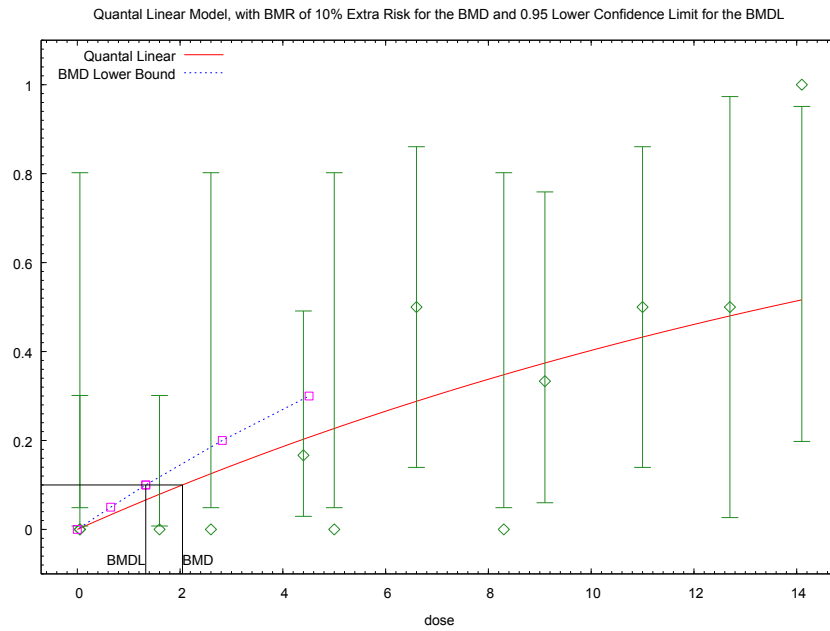
BMD = 3.97348

BMDL = 1.83708

BMDU = 5.1094

Taken together, (1.83708, 5.1094) is a 90 % two-sided confidence interval for the BMD

Cancer Slope Factor = 0.0544342



20:30 12/11 2016

```

=====
Quantal Linear Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_williams_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_williams_Opt.plt
Sun Dec 11 20:30:59 2016
=====

```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose})]$$

Dependent variable = Effect
Independent variable = Dose

Total number of observations = 12
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 (and Specified) Parameter Values
Background =      0.125
Slope =         0.0888484
Power =          1   Specified

```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Power
have been estimated at a boundary point, or have been specified by
the user,
and do not appear in the correlation matrix)
Slope

Slope 1

Parameter Estimates

95.0% Wald Confidence

Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper Conf. Limit
	Background	0	NA		
	Slope	0.0514667	0.0144263	0.0231916	
0.0797417					

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-18.9299	12			
Fitted model	-23.6376	1	9.4154	11	0.5836
Reduced model	-32.7474	1	27.635	11	0.003681
AIC:	49.2752				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0500	0.0026	0.031	0.000	12.000	-0.176
1.6000	0.0790	0.949	0.000	12.000	-1.015
4.4000	0.2026	2.432	2.000	12.000	-0.310
6.6000	0.2880	1.728	3.000	6.000	1.147
9.1000	0.3740	2.244	2.000	6.000	-0.206
11.0000	0.4323	2.594	3.000	6.000	0.335
0.0500	0.0026	0.005	0.000	2.000	-0.072
2.6000	0.1252	0.250	0.000	2.000	-0.535
5.0000	0.2269	0.454	0.000	2.000	-0.766
8.3000	0.3476	0.695	0.000	2.000	-1.032
12.7000	0.4798	0.960	1.000	2.000	0.057
14.1000	0.5160	1.032	2.000	2.000	1.370

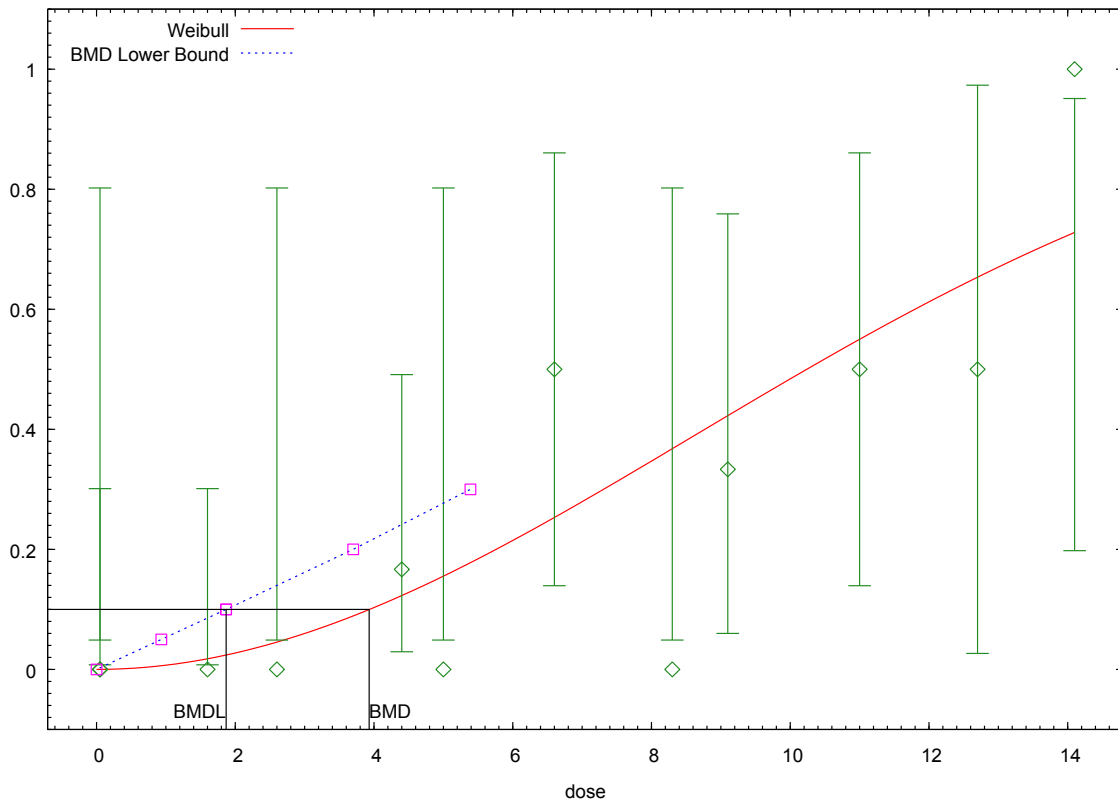
Chi^2 = 6.45 d.f. = 11 P-value = 0.8417

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 2.04716
 BMDL = 1.33219

Output of Weibull model not shown.

Weibull Model, with BMR of 10% Extra Risk for the BMD and 0.95 Lower Confidence Limit for the BMDL



09:25 06/28 2017

```

=====
      Weibull Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_williams_Opt.(d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_williams_Opt.plt
                                          Wed Jun 28 09:25:29 2017
=====
    
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose}^{\text{power}})]$$

Dependent variable = Effect
 Independent variable = Dose
 Power parameter is not restricted

Total number of observations = 12
 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 (and Specified) Parameter Values
Background =      0.125
Slope =         0.0460466
Power =         1.24839
    
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	Slope	Power
Slope	1	-0.98
Power	-0.98	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
0.0275626	Slope	0.00708737	0.0104468	-0.0133879	
3.29451	Power	1.97194	0.674791	0.649375	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-18.9299	12			
Fitted model	-22.2922	2	6.72465	10	0.7512
Reduced model	-32.7474	1	27.635	11	0.003681

AIC: 48.5844

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0500	0.0000	0.000	0.000	12.000	-0.015
1.6000	0.0177	0.213	0.000	12.000	-0.466
4.4000	0.1233	1.480	2.000	12.000	0.457
6.6000	0.2538	1.523	3.000	6.000	1.386
9.1000	0.4240	2.544	2.000	6.000	-0.449
11.0000	0.5515	3.309	3.000	6.000	-0.253
0.0500	0.0000	0.000	0.000	2.000	-0.006
2.6000	0.0456	0.091	0.000	2.000	-0.309
5.0000	0.1558	0.312	0.000	2.000	-0.608
8.3000	0.3688	0.738	0.000	2.000	-1.081
12.7000	0.6551	1.310	1.000	2.000	-0.461
14.1000	0.7297	1.459	2.000	2.000	0.861

Chi^2 = 5.20 d.f. = 10 P-value = 0.8775

Benchmark Dose Computation

Specified effect = 0.1

Risk Type = Extra risk
Confidence level = 0.95
BMD = 3.9304
BMDL = 1.86814

Supplementary information for DOGSVOMIT (Hughes et al., 1999)

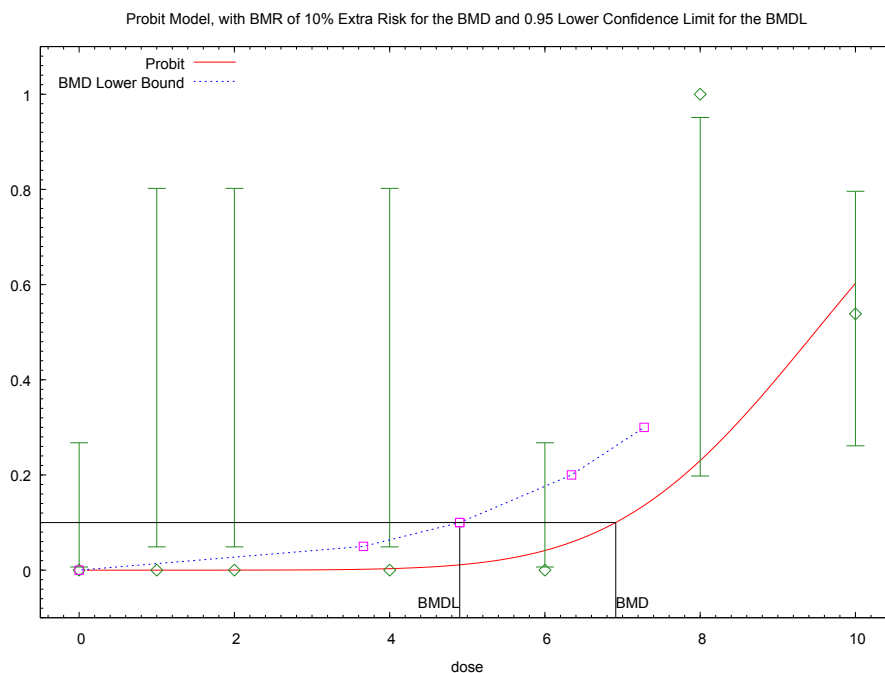
Details of the BMD Analysis of concentration-response data for vomiting in dogs of Hughes et al. (1999) reported in Appendix G, Section G.4.2.1 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

Filename: DogsVomiting Folder: HughesVomiting

Output from BMDS from 19 Oct 2016 with the full information of the fit of all models of BMDS where a model fit was obtained in a reasonable duration of computing time.

The output is ordered in the sequence used in the Table G14 shown in Appendix G which is based on an order of increasing mathematical complexity. The two less complex multistage models are nested under the multistage model which was used with the default assumption of the degree of the polynomial of BMDS.

Occasionally, it was not possible to obtain graphic of the Gamma model with the notebook used for the calculation HP8440p or Lenovo 3508.



19:25 10/19 2016

```

=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Hugehsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Hugehsvomit_Opt.plt
Wed Oct 19 19:25:22 2016
=====

```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Dose}),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 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 (and Specified) Parameter Values

background = 0 Specified
 intercept = -2.44048
 slope = 0.212598

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-0.98
slope	-0.98	1

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence	
				Lower Conf. Limit	Upper
Conf. Limit	intercept	-4.72924	1.59684	-7.85898	-
1.59949	slope	0.498785	0.175706	0.154408	
0.843161					

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-12.6161	2	7.28735	5	0.2001
Reduced model	-23.369	1	28.7932	6	<.0001
AIC:	29.2322				

Goodness of Fit

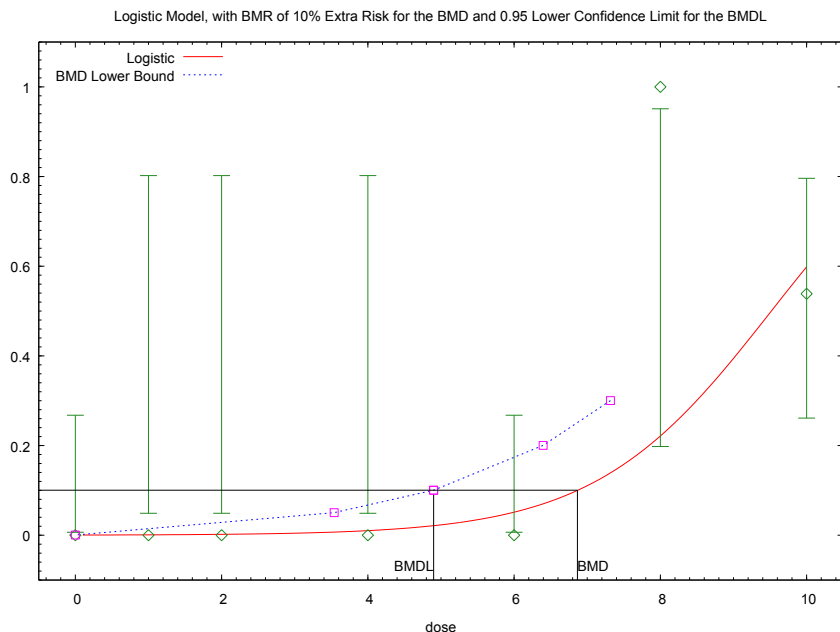
Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	14.000	-0.004
1.0000	0.0000	0.000	0.000	2.000	-0.005
2.0000	0.0001	0.000	0.000	2.000	-0.014
4.0000	0.0031	0.006	0.000	2.000	-0.079
6.0000	0.0412	0.577	0.000	14.000	-0.776
8.0000	0.2300	0.460	2.000	2.000	2.588

10.0000 0.6020 7.826 7.000 13.000 -0.468
 Chi² = 7.52 d.f. = 5 P-value = 0.1844

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95

 BMD = 6.91218
 BMDL = 4.90252



19:27 10/19 2016

```
=====
Logistic Model. (Version: 2.14; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Hugehsgdogsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Hugehsgdogsvomit_Opt.plt
Wed Oct 19 19:27:55 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = 1/[1+\text{EXP}(-\text{intercept}-\text{slope}*\text{dose})]$$

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 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
background = 0 Specified
intercept = -2.79186
slope = 0.313956

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background
have been estimated at a boundary point, or have been specified by
the user,
and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-0.98
slope	-0.98	1

Parameter Estimates

Interval Variable Conf. Limit	Estimate	Std. Err.	95.0% Wald Confidence	
			Lower Conf. Limit	Upper
intercept 2.36055	-7.88813	2.82024	-13.4157	-
slope 1.4259	0.829423	0.304333	0.232942	

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-12.8551	2	7.76543	5	0.1696
Reduced model	-23.369	1	28.7932	6	<.0001
AIC:	29.7103				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0004	0.005	0.000	14.000	-0.072
1.0000	0.0009	0.002	0.000	2.000	-0.041
2.0000	0.0020	0.004	0.000	2.000	-0.063
4.0000	0.0102	0.020	0.000	2.000	-0.144
6.0000	0.0516	0.722	0.000	14.000	-0.873
8.0000	0.2222	0.444	2.000	2.000	2.646
10.0000	0.6002	7.802	7.000	13.000	-0.454

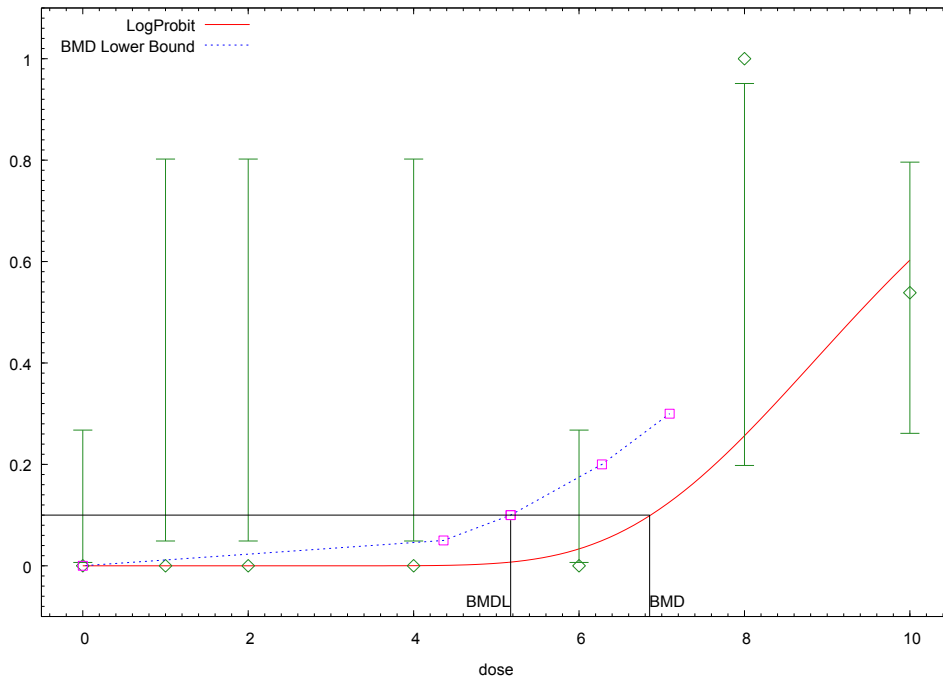
Chi^2 = 8.00 d.f. = 5 P-value = 0.1563

Benchmark Dose Computation

Specified effect = 0.1
Risk Type = Extra risk

Confidence level = 0.95
 BMD = 6.86579
 BMDL = 4.89953

LogProbit Model, with BMR of 10% Extra Risk for the BMD and 0.95 Lower Confidence Limit for the BMDL



19:29 10/19 2016

```
=====  

  Probit Model. (Version: 3.3; Date: 2/28/2013)  

  Input Data File:  

  C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Hugehsvomit_Opt.(d)  

  Gnuplot Plotting File:  

  C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Hugehsvomit_Opt.plt  

  Wed Oct 19 19:29:07 2016  

  =====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{Background} + (1 - \text{Background}) * \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Log}(\text{Dose})),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 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

User has chosen the log transformed model

Default Initial (and Specified) Parameter Values
 background = 0
 intercept = -1.95857
 slope = 0.635671

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-1
slope	-1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	background	0	NA		
2.80386	intercept	-9.17682	3.25157	-15.5498	-
7.00916	slope	4.10195	1.4833	1.19475	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-12.2795	2	6.61423	5	0.2509
Reduced model	-23.369	1	28.7932	6	<.0001
AIC:	28.5591				

Goodness of Fit

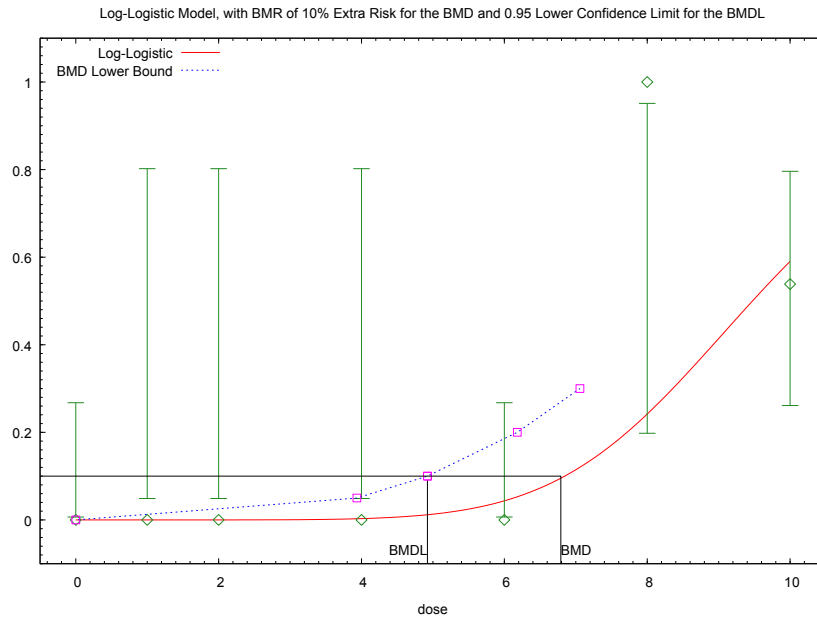
Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	14.000	0.000
1.0000	0.0000	0.000	0.000	2.000	-0.000
2.0000	0.0000	0.000	0.000	2.000	-0.000
4.0000	0.0002	0.000	0.000	2.000	-0.022
6.0000	0.0338	0.474	0.000	14.000	-0.700
8.0000	0.2588	0.518	2.000	2.000	2.393
10.0000	0.6058	7.875	7.000	13.000	-0.497

Chi^2 = 6.47 d.f. = 5 P-value = 0.2635

Benchmark Dose Computation

Specified effect = 0.1

Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 6.85349
 BMDL = 5.1729



19:30 10/19 2016

```
=====
Logistic Model. (Version: 2.14; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Hugehsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Hugehsvomit_Opt.plt
Wed Oct 19 19:30:34 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 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

User has chosen the log transformed model

Default Initial Parameter Values
 background = 0
 intercept = -2.19376

slope = 0.815501

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-1
slope	-1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	background	0	NA		
3.74686	intercept	-15.1551	5.82063	-26.5633	-
11.9209	slope	6.76452	2.63088	1.60809	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-12.5076	2	7.07046	5	0.2155
Reduced model	-23.369	1	28.7932	6	<.0001
AIC:	29.0153				

Goodness of Fit

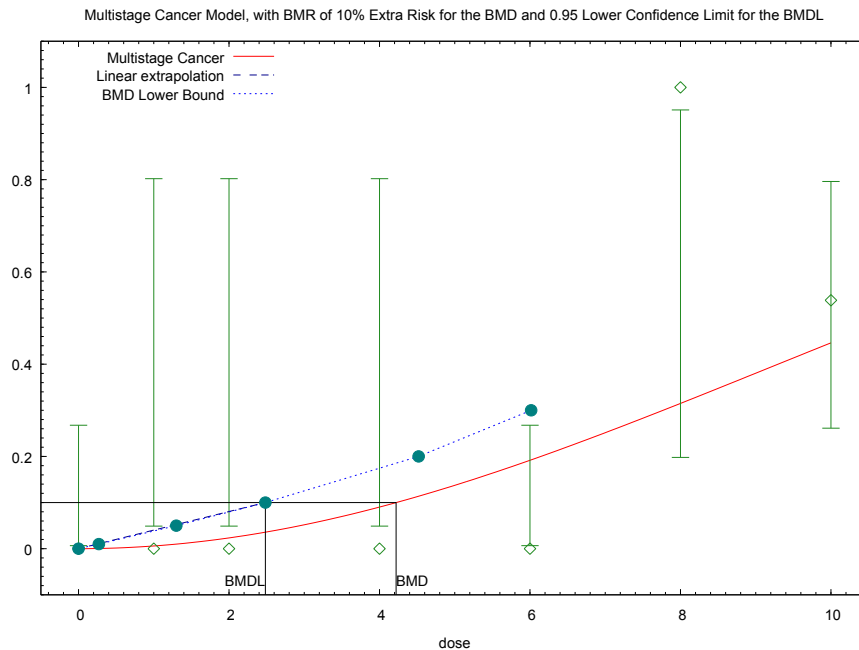
Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	14.000	0.000
1.0000	0.0000	0.000	0.000	2.000	-0.001
2.0000	0.0000	0.000	0.000	2.000	-0.008
4.0000	0.0031	0.006	0.000	2.000	-0.079
6.0000	0.0459	0.642	0.000	14.000	-0.821
8.0000	0.2519	0.504	2.000	2.000	2.437
10.0000	0.6037	7.848	7.000	13.000	-0.481

Chi^2 = 6.85 d.f. = 5 P-value = 0.2319

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95

BMD = 6.79077
 BMDL = 4.92267



19:32 10/19 2016

```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Hugehsvommit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Hugehsvommit_Opt.plt
Wed Oct 19 19:32:07 2016
=====
    
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1 - \text{EXP}(-\text{beta1} * \text{dose}^1 - \text{beta2} * \text{dose}^2)]$$

The parameter betas are restricted to be positive

Dependent variable = Effect
 Independent variable = Dose

Total number of observations = 7
 Total number of records with missing values = 0
 Total number of parameters in model = 3
 Total number of specified parameters = 0
 Degree of polynomial = 2

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

Background = 0
 Beta(1) = 4.26621e+018
 Beta(2) = 0

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Beta(1) have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

Beta(2)
 Beta(2) 1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
	Beta(1)	0	NA		
	Beta(2)	0.00591458	0.00199656	0.0020014	
		0.00982776			

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-14.7321	1	11.5194	6	0.07359
Reduced model	-23.369	1	28.7932	6	<.0001
AIC:	31.4643				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	14.000	-0.000
1.0000	0.0059	0.012	0.000	2.000	-0.109
2.0000	0.0234	0.047	0.000	2.000	-0.219
4.0000	0.0903	0.181	0.000	2.000	-0.446
6.0000	0.1918	2.685	0.000	14.000	-1.823
8.0000	0.3151	0.630	2.000	2.000	2.085
10.0000	0.4465	5.804	7.000	13.000	0.667

Chi^2 = 8.37 d.f. = 6 P-value = 0.2121

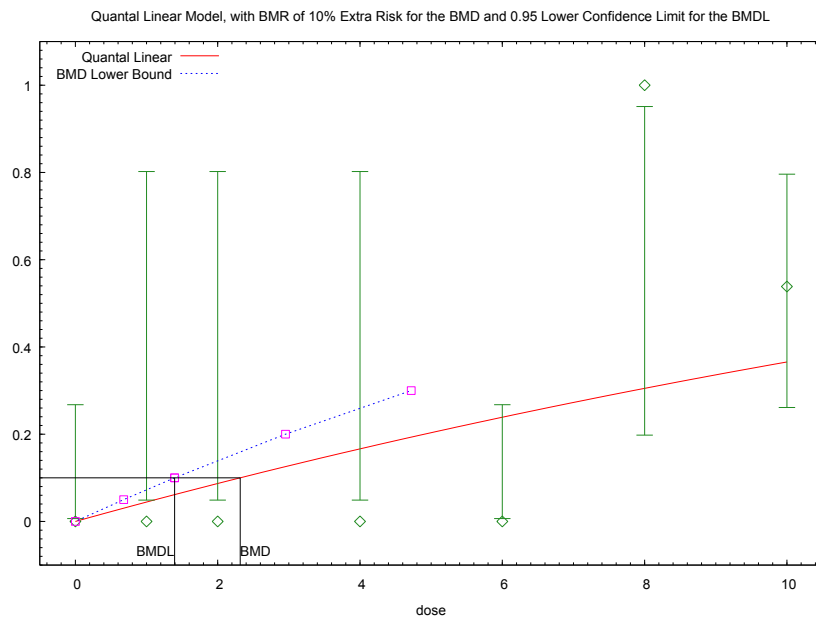
Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95

BMD = 4.22063
 BMDL = 2.48314
 BMDU = 5.72306

Taken together, (2.48314, 5.72306) is a 90 % two-sided confidence interval for the BMD

Cancer Slope Factor = 0.0402716



19:33 10/19 2016

```
=====
Quantal Linear Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Hugehsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Hugehsvomit_Opt.plt
Wed Oct 19 19:33:30 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose})]$$

Dependent variable = Effect
 Independent variable = Dose

Total number of observations = 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 (and Specified) Parameter Values

Background = 0.0625
 Slope = 0.0697602
 Power = 1 Specified

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Power
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

Slope

Slope	1
-------	---

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
0.0754358	Slope	0.045484	0.0152818	0.0155322	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-16.6076	1	15.2704	6	0.01825
Reduced model	-23.369	1	28.7932	6	<.0001

AIC: 35.2152

Goodness of Fit

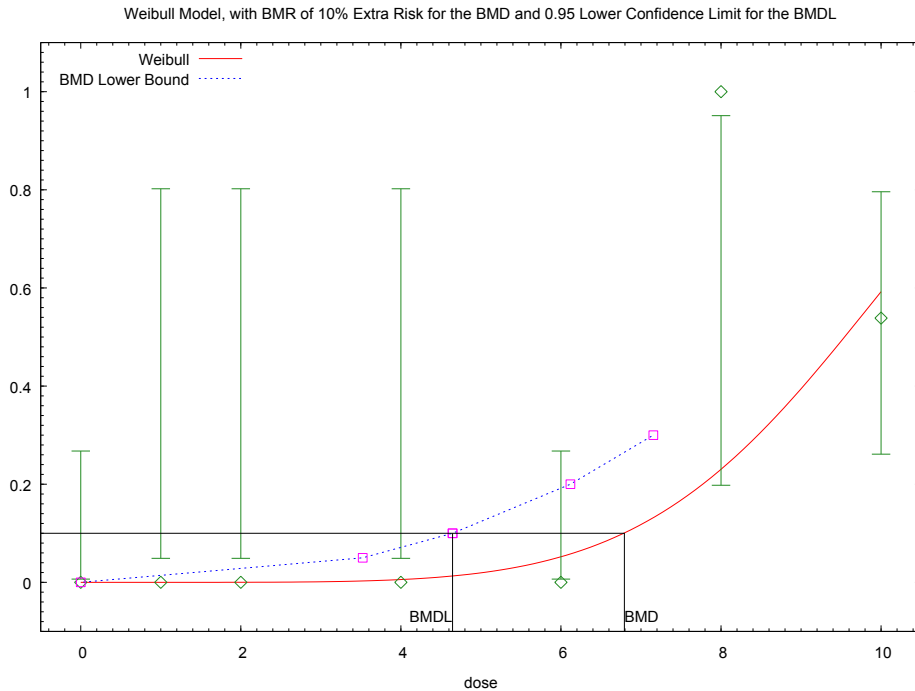
Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	14.000	0.000
1.0000	0.0445	0.089	0.000	2.000	-0.305
2.0000	0.0870	0.174	0.000	2.000	-0.436
4.0000	0.1663	0.333	0.000	2.000	-0.632
6.0000	0.2388	3.344	0.000	14.000	-2.096
8.0000	0.3050	0.610	2.000	2.000	2.135
10.0000	0.3655	4.751	7.000	13.000	1.295

Chi^2 = 11.31 d.f. = 6 P-value = 0.0792

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95

BMD = 2.31643
 BMDL = 1.39429



19:34 10/19 2016

```
=====
Weibull Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Hugehsgdogsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Hugehsgdogsvomit_Opt.plt
Wed Oct 19 19:34:50 2016
=====
```

```
BMDS_Model_Run
~~~~~
```

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose}^{\text{power}})]$$

Dependent variable = Effect
 Independent variable = Dose
 Power parameter is not restricted

Total number of observations = 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 (and Specified) Parameter Values
Background = 0.0625
Slope = 0.00453905
Power = 2.18664
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	Slope	Power
Slope	1	-1
Power	-1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper Conf. Limit
	Background	0	NA		
2.9962e-005	Slope	2.70866e-006	1.3905e-005	-2.45447e-005	
9.97704	Power	5.51646	2.27585	1.05589	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-12.7463	2	7.54776	5	0.183
Reduced model	-23.369	1	28.7932	6	<.0001
AIC:	29.4926				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	14.000	0.000
1.0000	0.0000	0.000	0.000	2.000	-0.002
2.0000	0.0001	0.000	0.000	2.000	-0.016
4.0000	0.0057	0.011	0.000	2.000	-0.107
6.0000	0.0517	0.724	0.000	14.000	-0.874
8.0000	0.2288	0.458	2.000	2.000	2.597
10.0000	0.5892	7.660	7.000	13.000	-0.372

Chi^2 = 7.66 d.f. = 5 P-value = 0.1763

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 6.79268

BMDL = 4.64587

Gamma dogs

```
=====
      Gamma Model. (Version: 2.16; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/gam_Hugehsgdogsvomit_Opt.(d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/gam_Hugehsgdogsvomit_Opt.plt
                                          Thu Oct 27 11:29:08 2016
=====
```

```
BMDS_Model_Run
~~~~~
```

The form of the probability function is:

$P[\text{response}] = \text{background} + (1 - \text{background}) * \text{CumGamma}[\text{slope} * \text{dose}, \text{power}]$,
 where CumGamma(.) is the cumulative Gamma distribution function

Dependent variable = Effect
 Independent variable = Dose
 Power parameter is not restricted

Total number of observations = 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 (and Specified) Parameter Values
Background = 0.0625
Slope = 0.0710638
Power = 1.3
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Power
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

```
Slope
Slope      1
```

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence	
				Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
2.13317	Slope	1.87644	0.13099	1.6197	

Power 18 NA

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-8.97241	7			
Fitted model	-12.388	1	6.83117	6	0.3367
Reduced model	-23.369	1	28.7932	6	<.0001

AIC: 26.776

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	14.000	0.000
1.0000	0.0000	0.000	0.000	2.000	-0.000
2.0000	0.0000	0.000	0.000	2.000	-0.000
4.0000	0.0008	0.002	0.000	2.000	-0.040
6.0000	0.0388	0.543	0.000	14.000	-0.751
8.0000	0.2521	0.504	2.000	2.000	2.436
10.0000	0.6011	7.814	7.000	13.000	-0.461

Chi² = 6.71 d.f. = 6 P-value = 0.3484

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 6.83297
 BMDL = 5.05101

Supplementary information for CATSVOMIT (Hughes et al., 1999)

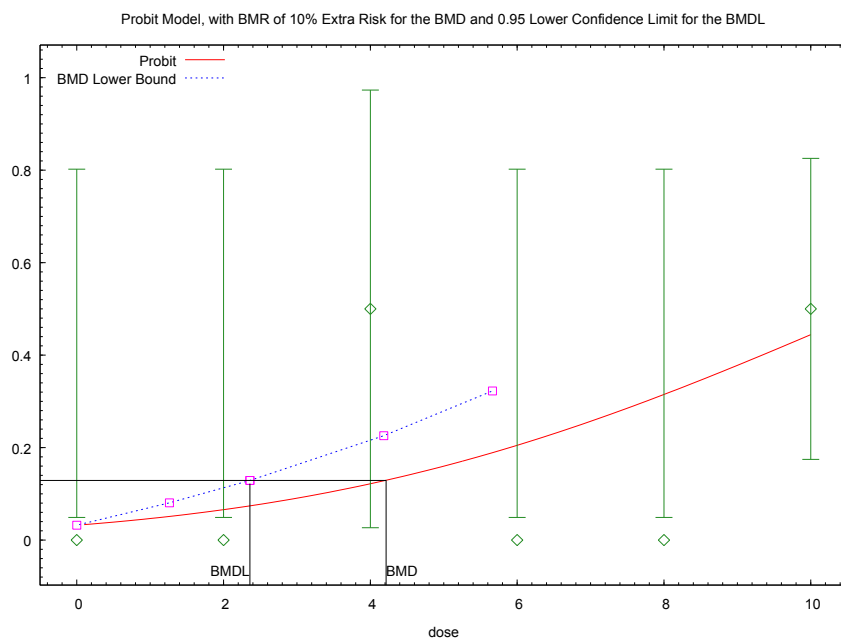
Details of the BMD Analysis of concentration-response data for vomiting in dogs of Hughes et al. (1999) reported in Appendix G, Section G.4.2.2 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed

Filename: CatsVomiting Folder: HughesVomiting

Output from BMDS from 19 Oct 2016 with the full information of the fit of all models of BMDS where a model fit was obtained in a reasonable duration of computing time.

The output is ordered in the sequence used in the Table G15 shown in Appendix G which is based on an order of increasing mathematical complexity. The two less complex multistage models are nested under the multistage model which was used with the default assumption of the degree of the polynomial of BMDS.

Occasionally, it was not possible to obtain graphic of the Gamma model with the notebook used for the calculation HP8440p or Lenovo 3508.



19:41 10/19 2016

```

=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Hugehscatsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Hugehscatsvomit_Opt.plt
Wed Oct 19 19:41:07 2016
=====

```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Dose}),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect

Independent variable = Dose

Slope parameter is not restricted

Total number of observations = 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 (and Specified) Parameter Values
 background = 0 Specified
 intercept = -1.49287
 slope = 0.12796

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-0.94
slope	-0.94	1

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence	
				Lower Conf. Limit	Upper
Conf. Limit	intercept	-1.85036	0.962378	-3.73659	
0.0358636	slope	0.17058	0.114444	-0.053725	
0.394886					

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-6.93147	6			
Fitted model	-9.2468	2	4.63066	4	0.3273
Reduced model	-10.6352	1	7.40738	5	0.1921
AIC:	22.4936				

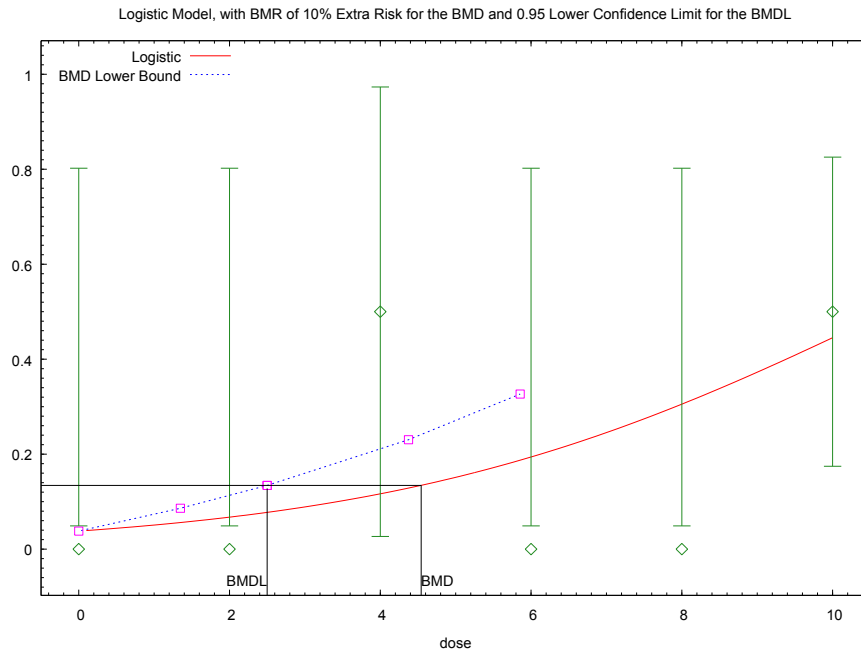
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0321	0.064	0.000	2.000	-0.258
2.0000	0.0656	0.131	0.000	2.000	-0.375
4.0000	0.1214	0.243	1.000	2.000	1.639
6.0000	0.2042	0.408	0.000	2.000	-0.716
8.0000	0.3136	0.627	0.000	2.000	-0.956
10.0000	0.4425	3.540	4.000	8.000	0.327

Chi^2 = 4.43 d.f. = 4 P-value = 0.3511

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 4.21409
 BMDL = 2.35852



19:42 10/19 2016

```
=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Hugehscatsvomit_Opt.(d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Hugehscatsvomit_Opt.plt
                                     Wed Oct 19 19:42:20 2016
=====
```

BMDS_Model_Run

~~~~~

The form of the probability function is:

$$P[\text{response}] = 1/[1+\text{EXP}(-\text{intercept}-\text{slope}*\text{dose})]$$

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 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  
background = 0 Specified  
intercept = -1.5328  
slope = 0.0919679

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
have been estimated at a boundary point, or have been specified by  
the user,  
and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.95 |
| slope     | -0.95     | 1     |

Parameter Estimates

| Interval    | Variable  | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|-----------|----------|-----------|-----------------------|-------|
|             |           |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | intercept | -3.23272 | 1.90545   | -6.96734              |       |
| 0.501899    | slope     | 0.301159 | 0.217233  | -0.124611             |       |
| 0.726928    |           |          |           |                       |       |

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -6.93147        | 6         |          |           |         |
| Fitted model  | -9.24436        | 2         | 4.62577  | 4         | 0.3279  |
| Reduced model | -10.6352        | 1         | 7.40738  | 5         | 0.1921  |
| AIC:          | 22.4887         |           |          |           |         |

Goodness of Fit

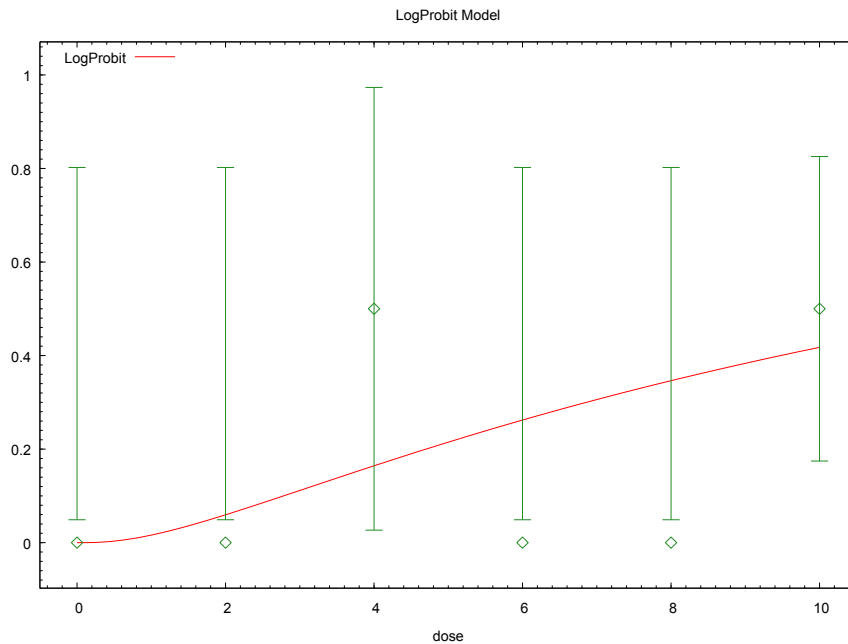
| Dose    | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|---------|------------|----------|----------|-------|-----------------|
| 0.0000  | 0.0380     | 0.076    | 0.000    | 2.000 | -0.281          |
| 2.0000  | 0.0672     | 0.134    | 0.000    | 2.000 | -0.380          |
| 4.0000  | 0.1163     | 0.233    | 1.000    | 2.000 | 1.693           |
| 6.0000  | 0.1938     | 0.388    | 0.000    | 2.000 | -0.693          |
| 8.0000  | 0.3050     | 0.610    | 0.000    | 2.000 | -0.937          |
| 10.0000 | 0.4449     | 3.560    | 4.000    | 8.000 | 0.313           |

Chi^2 = 4.55      d.f. = 4      P-value = 0.3372

Benchmark Dose Computation

Specified effect = 0.1  
Risk Type = Extra risk  
Confidence level = 0.95  
BMD = 4.54256

BMDL = 2.49929



19:44 10/19 2016

```

=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Hugehscatsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Hugehscatsvomit_Opt.plt
Wed Oct 19 19:44:39 2016
=====
    
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{Background} + (1 - \text{Background}) * \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Log}(\text{Dose})),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 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

User has chosen the log transformed model

```

Default Initial (and Specified) Parameter Values
background = 0
intercept = -1.21474
slope = 0.44955
    
```

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|           |           |       |
|-----------|-----------|-------|
|           | intercept | slope |
| intercept | 1         | -0.98 |
| slope     | -0.98     | 1     |

Parameter Estimates

|             |            | 95.0% Wald Confidence |           |                   |       |
|-------------|------------|-----------------------|-----------|-------------------|-------|
| Interval    | Variable   | Estimate              | Std. Err. | Lower Conf. Limit | Upper |
| Conf. Limit | background | 0                     | NA        |                   |       |
| 0.983281    | intercept  | -2.1403               | 1.59369   | -5.26388          |       |
| 2.35276     | slope      | 0.839456              | 0.772106  | -0.673844         |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -6.93147        | 6         |          |           |         |
| Fitted model  | -9.22197        | 2         | 4.581    | 4         | 0.3331  |
| Reduced model | -10.6352        | 1         | 7.40738  | 5         | 0.1921  |
| AIC:          | 22.4439         |           |          |           |         |

Goodness of Fit

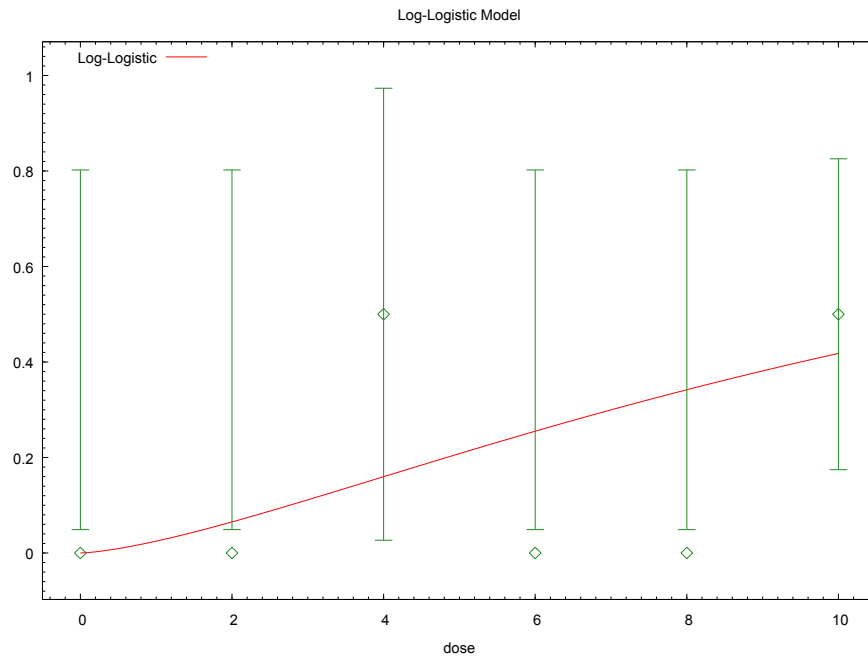
| Dose    | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|---------|------------|----------|----------|-------|-----------------|
| 0.0000  | 0.0000     | 0.000    | 0.000    | 2.000 | 0.000           |
| 2.0000  | 0.0596     | 0.119    | 0.000    | 2.000 | -0.356          |
| 4.0000  | 0.1644     | 0.329    | 1.000    | 2.000 | 1.281           |
| 6.0000  | 0.2623     | 0.525    | 0.000    | 2.000 | -0.843          |
| 8.0000  | 0.3465     | 0.693    | 0.000    | 2.000 | -1.030          |
| 10.0000 | 0.4179     | 3.343    | 4.000    | 8.000 | 0.471           |

Chi^2 = 3.76      d.f. = 4      P-value = 0.4394

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 2.78147

Benchmark dose computation failed. Lower limit



19:45 10/19 2016

```

=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Hugehscatsvomit_Opt.(d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Hugehscatsvomit_Opt.plt
                                          Wed Oct 19 19:45:46 2016
=====
  
```

```

BMDS_Model_Run
~~~~~

```

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 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

User has chosen the log transformed model

Default Initial Parameter Values  
 background = 0

intercept = -1.60304  
 slope = 0.386137

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -0.98 |
| slope     | -0.98     | 1     |

Parameter Estimates

| Interval    | Variable   | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|----------|-----------|-----------------------|-------|
|             |            |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | background | 0        | NA        |                       |       |
| 2.22047     | intercept  | -3.67491 | 3.0079    | -9.57029              |       |
| 4.25207     | slope      | 1.45434  | 1.42744   | -1.34338              |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -6.93147        | 6         |          |           |         |
| Fitted model  | -9.22265        | 2         | 4.58236  | 4         | 0.3329  |
| Reduced model | -10.6352        | 1         | 7.40738  | 5         | 0.1921  |
| AIC:          | 22.4453         |           |          |           |         |

Goodness of Fit

| Dose    | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|---------|------------|----------|----------|-------|-----------------|
| 0.0000  | 0.0000     | 0.000    | 0.000    | 2.000 | 0.000           |
| 2.0000  | 0.0650     | 0.130    | 0.000    | 2.000 | -0.373          |
| 4.0000  | 0.1599     | 0.320    | 1.000    | 2.000 | 1.312           |
| 6.0000  | 0.2556     | 0.511    | 0.000    | 2.000 | -0.829          |
| 8.0000  | 0.3428     | 0.686    | 0.000    | 2.000 | -1.021          |
| 10.0000 | 0.4192     | 3.353    | 4.000    | 8.000 | 0.463           |

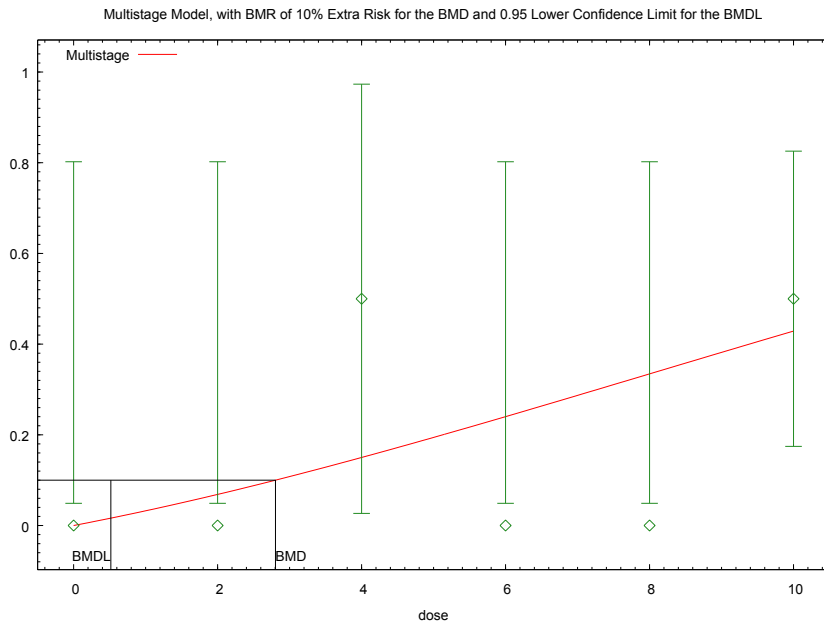
Chi^2 = 3.81      d.f. = 4      P-value = 0.4330

BMDL curve computation failed for BMR = 0.050000 .  
 The BMDL curve appearing in the graph may not be accurate.  
 BMDL curve computation failed for BMR = 0.100000 .  
 The BMDL curve appearing in the graph may not be accurate.  
 BMDL curve computation failed for BMR = 0.200000 .  
 The BMDL curve appearing in the graph may not be accurate.  
 BMDL curve computation failed for BMR = 0.300000 .  
 The BMDL curve appearing in the graph may not be accurate.



Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 2.76226



19:47 10/19 2016

Benchmark dose computation failed. Lower li

```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Hugehscatsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Hugehscatsvomit_Opt.plt
Wed Oct 19 19:47:07 2016
=====

```

BMDS\_Model\_Run

~~~~~

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1 - \text{EXP}(-\text{beta1} * \text{dose}^{1-\text{beta2}} * \text{dose}^2)]$$

The parameter betas are not restricted

Dependent variable = Effect  
 Independent variable = Dose

Total number of observations = 6  
 Total number of records with missing values = 0  
 Total number of parameters in model = 3

Total number of specified parameters = 0  
 Degree of polynomial = 2

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

Background = 0.0715751  
 Beta(1) = 0.00866434  
 Beta(2) = 0.00309441

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|         | Beta(1) | Beta(2) |
|---------|---------|---------|
| Beta(1) | 1       | -0.94   |
| Beta(2) | -0.94   | 1       |

Parameter Estimates

| Interval    | Variable   | Estimate  | Std. Err.  | 95.0% Wald Confidence |       |
|-------------|------------|-----------|------------|-----------------------|-------|
| Conf. Limit |            |           |            | Lower Conf. Limit     | Upper |
|             | Background | 0         | NA         |                       |       |
| 0.168342    | Beta(1)    | 0.0304085 | 0.0703757  | -0.107525             |       |
| 0.0187505   | Beta(2)    | 0.0025553 | 0.00826301 | -0.0136399            |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -6.93147        | 6         |          |           |         |
| Fitted model  | -9.19187        | 2         | 4.5208   | 4         | 0.3401  |
| Reduced model | -10.6352        | 1         | 7.40738  | 5         | 0.1921  |
| AIC:          | 22.3837         |           |          |           |         |

Goodness of Fit

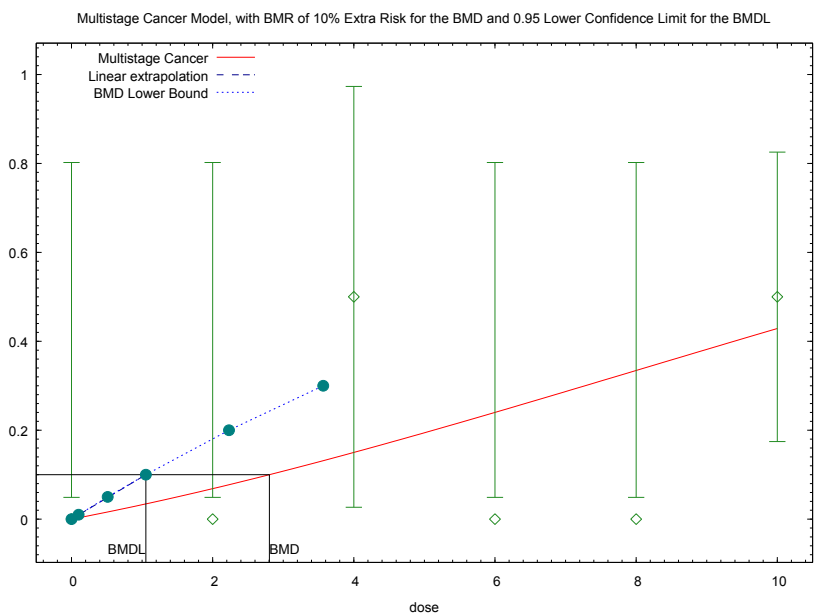
| Dose   | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|--------|------------|----------|----------|-------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 2.000 | 0.000           |
| 2.0000 | 0.0686     | 0.137    | 0.000    | 2.000 | -0.384          |
| 4.0000 | 0.1500     | 0.300    | 1.000    | 2.000 | 1.386           |
| 6.0000 | 0.2400     | 0.480    | 0.000    | 2.000 | -0.795          |
| 8.0000 | 0.3342     | 0.668    | 0.000    | 2.000 | -1.002          |

10.0000      0.4286            3.429      4.000            8.000            0.408  
 Chi^2 = 3.87      d.f. = 4            P-value = 0.4237

Benchmark Dose Computation

Specified effect =            0.1  
 Risk Type            =            Extra risk  
 Confidence level =            0.95  
  
                           BMD =            2.80409  
                           BMDL =            0.517005

BMDU did not converge for BMR = 0.100000  
 BMDU calculation failed  
                           BMDU =            2.80409



19:48 10/19 2016

```
=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Hugehscatsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Hugehscatsvomit_Opt.plt
Wed Oct 19 19:48:54 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:  

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1 - \text{EXP}(-\text{beta1} * \text{dose}^{1-\text{beta2}} * \text{dose}^2)]$$

The parameter betas are restricted to be positive

Dependent variable = Effect  
 Independent variable = Dose

Total number of observations = 6  
 Total number of records with missing values = 0  
 Total number of parameters in model = 3  
 Total number of specified parameters = 0  
 Degree of polynomial = 2

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

Background = 0.0715751  
 Beta(1) = 0.00866434  
 Beta(2) = 0.00309441

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|         | Beta(1) | Beta(2) |
|---------|---------|---------|
| Beta(1) | 1       | -0.94   |
| Beta(2) | -0.94   | 1       |

Parameter Estimates

| Interval<br>Variable<br>Conf. Limit | Estimate  | Std. Err.  | 95.0% Wald Confidence |       |
|-------------------------------------|-----------|------------|-----------------------|-------|
|                                     |           |            | Lower                 | Upper |
| Background                          | 0         | NA         |                       |       |
| Beta(1)<br>0.168342                 | 0.0304085 | 0.0703757  | -0.107525             |       |
| Beta(2)<br>0.0187505                | 0.0025553 | 0.00826301 | -0.0136399            |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -6.93147        | 6         |          |           |         |
| Fitted model  | -9.19187        | 2         | 4.5208   | 4         | 0.3401  |
| Reduced model | -10.6352        | 1         | 7.40738  | 5         | 0.1921  |

AIC: 22.3837

Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|--------|------------|----------|----------|-------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 2.000 | 0.000           |

|         |        |       |       |       |        |
|---------|--------|-------|-------|-------|--------|
| 2.0000  | 0.0686 | 0.137 | 0.000 | 2.000 | -0.384 |
| 4.0000  | 0.1500 | 0.300 | 1.000 | 2.000 | 1.386  |
| 6.0000  | 0.2400 | 0.480 | 0.000 | 2.000 | -0.795 |
| 8.0000  | 0.3342 | 0.668 | 0.000 | 2.000 | -1.002 |
| 10.0000 | 0.4286 | 3.429 | 4.000 | 8.000 | 0.408  |

Chi<sup>2</sup> = 3.87      d.f. = 4      P-value = 0.4237

Benchmark Dose Computation

Specified effect = 0.1

Risk Type = Extra risk

Confidence level = 0.95

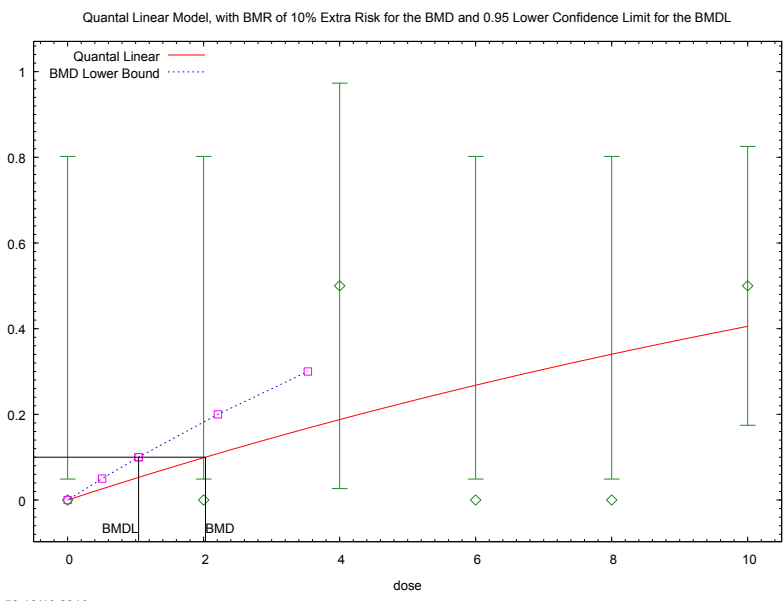
BMD = 2.80409

BMDL = 1.05353

BMDU = 44.3164

Taken together, (1.05353, 44.3164) is a 90 % two-sided confidence interval for the BMD

Cancer Slope Factor = 0.0949188



19:50 10/19 2016

```

=====
Quantal Linear Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Hugehscatsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Hugehscatsvomit_Opt.plt
Wed Oct 19 19:50:25 2016
=====

```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose})]$$

Dependent variable = Effect  
 Independent variable = Dose

Total number of observations = 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 (and Specified) Parameter Values  
 Background = 0.25  
 Slope = 0.0405465  
 Power = 1 Specified

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background -Power  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

Slope  
 Slope 1

Parameter Estimates

|             |            | 95.0% Wald Confidence |           |                   |       |
|-------------|------------|-----------------------|-----------|-------------------|-------|
| Interval    | Variable   | Estimate              | Std. Err. | Lower Conf. Limit | Upper |
| Conf. Limit | Background | 0                     | NA        |                   |       |
|             | Slope      | 0.0519717             | 0.0234603 | 0.00599037        |       |
|             |            |                       |           | 0.0979531         |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -6.93147        | 6         |          |           |         |
| Fitted model  | -9.23515        | 1         | 4.60737  | 5         | 0.4656  |
| Reduced model | -10.6352        | 1         | 7.40738  | 5         | 0.1921  |
| AIC:          | 20.4703         |           |          |           |         |

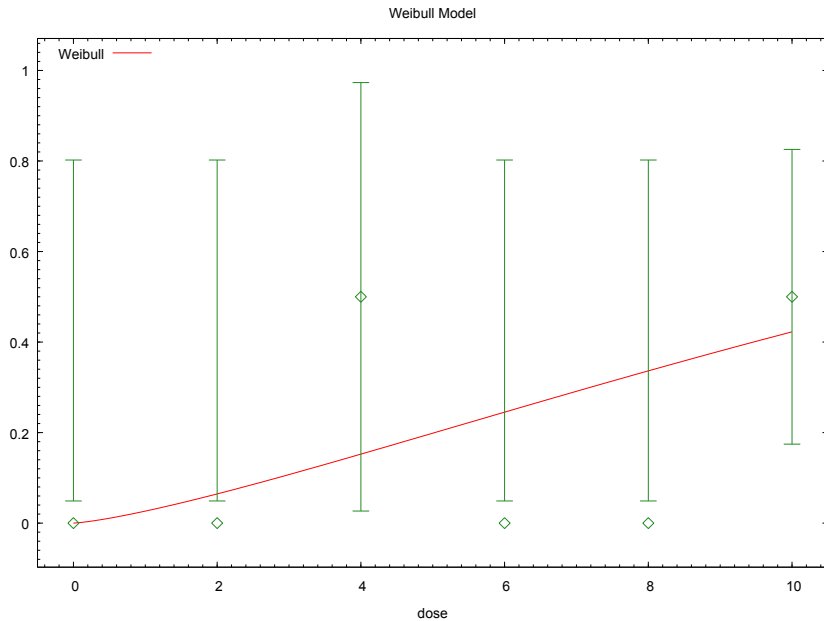
Goodness of Fit

| Dose    | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|---------|------------|----------|----------|-------|-----------------|
| 0.0000  | 0.0000     | 0.000    | 0.000    | 2.000 | 0.000           |
| 2.0000  | 0.0987     | 0.197    | 0.000    | 2.000 | -0.468          |
| 4.0000  | 0.1877     | 0.375    | 1.000    | 2.000 | 1.131           |
| 6.0000  | 0.2679     | 0.536    | 0.000    | 2.000 | -0.855          |
| 8.0000  | 0.3402     | 0.680    | 0.000    | 2.000 | -1.015          |
| 10.0000 | 0.4053     | 3.242    | 4.000    | 8.000 | 0.546           |

Chi^2 = 3.56      d.f. = 5      P-value = 0.6145

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 2.02727  
 BMDL = 1.04345



19:52 10/19 2016

```
=====
Weibull Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Hugehscatsvomit_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Hugehscatsvomit_Opt.plt
Wed Oct 19 19:52:16 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose}^{\text{power}})]$$

Dependent variable = Effect  
 Independent variable = Dose  
 Power parameter is not restricted

Total number of observations = 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 (and Specified) Parameter Values

Background = 0.25  
 Slope = 0.0879223  
 Power = 0.663855

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|       | Slope | Power |
|-------|-------|-------|
| Slope | 1     | -0.99 |
| Power | -0.99 | 1     |

Parameter Estimates

|             |            | 95.0% Wald Confidence |           |                   |       |
|-------------|------------|-----------------------|-----------|-------------------|-------|
| Interval    | Variable   | Estimate              | Std. Err. | Lower Conf. Limit | Upper |
| Conf. Limit | Background | 0                     | NA        |                   |       |
| 0.176706    | Slope      | 0.0269398             | 0.0764128 | -0.122827         |       |
| 3.9073      | Power      | 1.31346               | 1.32341   | -1.28037          |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -6.93147        | 6         |          |           |         |
| Fitted model  | -9.20446        | 2         | 4.54598  | 4         | 0.3371  |
| Reduced model | -10.6352        | 1         | 7.40738  | 5         | 0.1921  |
| AIC:          | 22.4089         |           |          |           |         |

Goodness of Fit

| Dose    | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|---------|------------|----------|----------|-------|-----------------|
| 0.0000  | 0.0000     | 0.000    | 0.000    | 2.000 | 0.000           |
| 2.0000  | 0.0648     | 0.130    | 0.000    | 2.000 | -0.372          |
| 4.0000  | 0.1533     | 0.307    | 1.000    | 2.000 | 1.361           |
| 6.0000  | 0.2468     | 0.494    | 0.000    | 2.000 | -0.810          |
| 8.0000  | 0.3387     | 0.677    | 0.000    | 2.000 | -1.012          |
| 10.0000 | 0.4256     | 3.405    | 4.000    | 8.000 | 0.426           |

Chi^2 = 3.85      d.f. = 4      P-value = 0.4265

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95



BMD = 2.82444

Benchmark dose computation failed. Lower limit i

## Supplementary information for WUDON (Wu et al., 2013a)

Details of the BMD Analysis of concentration-response data for vomiting in mink of Wu et al. (2013) reported in Appendix G, Section G.4.3.1 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed - for DON

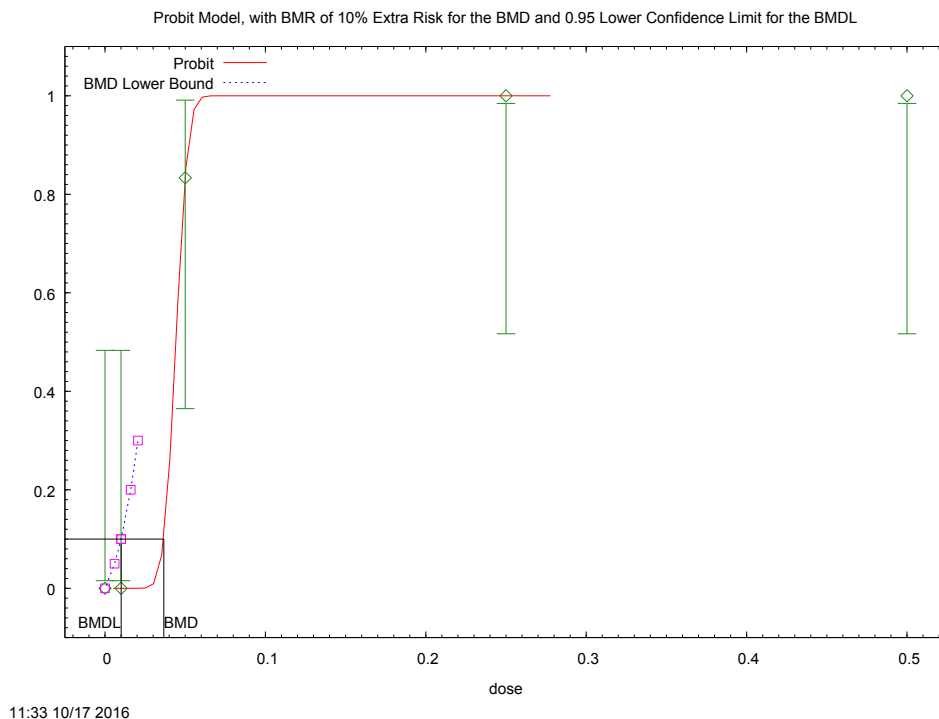
Filename: WuDON

Folder: Mink

Output from BMDS from 19 Oct 2016 with the full information of the fit of all models of BMDS where a model fit was obtained in a reasonable duration of computing time.

The output is ordered in the sequence used in the Table G16 shown in Appendix G which is based on an order of increasing mathematical complexity. The two less complex multistage models are nested under the multistage model which was used with the default assumption of the degree of the polynomial of BMDS.

Gamma model not shown below.



```

=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Wu2013DONb_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Wu2013DONb_Opt.plt
Mon Oct 17 11:33:07 2016
=====

```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Dose}),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 5  
 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 (and Specified) Parameter Values

background = 0 Specified  
 intercept = -0.977134  
 slope = 6.74251

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | -1    |
| slope     | -1        | 1     |

Parameter Estimates

| Interval    | Variable  | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|-----------|----------|-----------|-----------------------|-------|
|             |           |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | intercept | -7.46646 | 8633.95   | -16929.7              |       |
| 16914.8     | slope     | 168.677  | 172679    | -338276               |       |
| 338613      |           |          |           |                       |       |

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance     | Test d.f. | P-value |
|---------------|-----------------|-----------|--------------|-----------|---------|
| Full model    | -2.70337        | 5         |              |           |         |
| Fitted model  | -2.70337        | 2         | 4.52377e-008 | 3         | 1       |
| Reduced model | -20.527         | 1         | 35.6472      | 4         | <.0001  |
| AIC:          | 9.40673         |           |              |           |         |

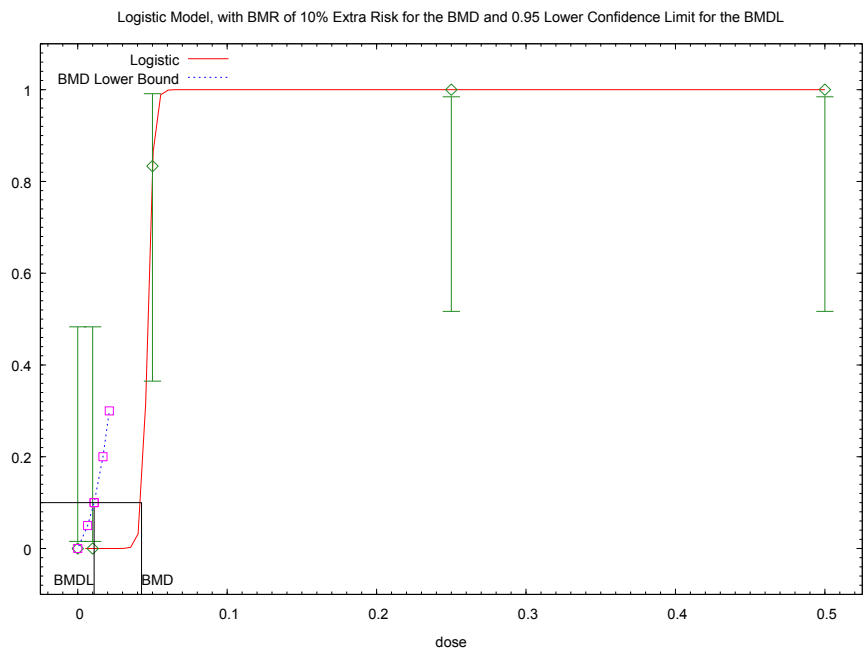
Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|--------|------------|----------|----------|-------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 6.000 | -0.000          |
| 0.0100 | 0.0000     | 0.000    | 0.000    | 6.000 | -0.000          |
| 0.0500 | 0.8333     | 5.000    | 5.000    | 6.000 | 0.000           |
| 0.2500 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |
| 0.5000 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |

Chi<sup>2</sup> = 0.00      d.f. = 3      P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 0.0366671  
 BMDL = 0.0100819



11:35 10/17 2016

```
=====
 Logistic Model. (Version: 2.14; Date: 2/28/2013)
 Input Data File:
 C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Wu2013DONb_Opt.(d)
 Gnuplot Plotting File:
 C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Wu2013DONb_Opt.plt
 Mon Oct 17 11:35:22 2016
=====
```

```
BMDS_Model_Run
~~~~~
```

The form of the probability function is:  
 $P[\text{response}] = 1/[1+\text{EXP}(-\text{intercept}-\text{slope}*\text{dose})]$

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted  
 Total number of observations = 5

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  
 background = 0 Specified  
 intercept = -1.28415  
 slope = 9.53091

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | NA        | NA    |
| slope     | NA        | NA    |

NA - This parameter's variance has been estimated as zero or less.  
 THE MODEL HAS PROBABLY NOT CONVERGED!!!

Parameter Estimates

|             |           | 95.0% Wald Confidence |           |                   |       |
|-------------|-----------|-----------------------|-----------|-------------------|-------|
| Interval    | Variable  | Estimate              | Std. Err. | Lower Conf. Limit | Upper |
| Conf. Limit | intercept | -24.4798              | NA        | NA                |       |
| NA          |           |                       |           |                   |       |
|             | slope     | 521.784               | NA        | NA                |       |
| NA          |           |                       |           |                   |       |

At least some variance estimates are negative.  
 THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!  
 Try again from another starting point.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance     | Test d.f. | P-value |
|---------------|-----------------|-----------|--------------|-----------|---------|
| Full model    | -2.70337        | 5         |              |           |         |
| Fitted model  | -2.70337        | 2         | 5.29225e-008 | 3         | 1       |
| Reduced model | -20.527         | 1         | 35.6472      | 4         | <.0001  |
| AIC:          | 9.40673         |           |              |           |         |

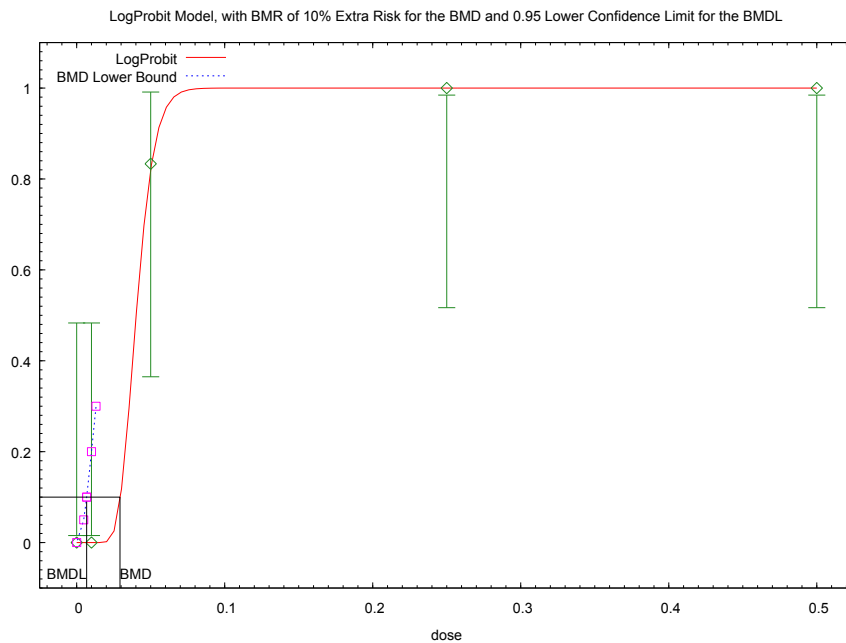
Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|--------|------------|----------|----------|-------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 6.000 | -0.000          |
| 0.0100 | 0.0000     | 0.000    | 0.000    | 6.000 | -0.000          |
| 0.0500 | 0.8333     | 5.000    | 5.000    | 6.000 | -0.000          |
| 0.2500 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |
| 0.5000 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |

Chi<sup>2</sup> = 0.00      d.f. = 3      P-value = 1.0000

Benchmark Dose Computation

Specified effect =                    0.1  
 Risk Type                    =        Extra risk  
 Confidence level =                    0.95  
                                           BMD =            0.0427045  
                                           BMDL =           0.0109621



11:36 10/17 2016

```
=====
      Probit Model. (Version: 3.3; Date: 2/28/2013)
      Input Data File:
      C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Wu2013DONb_Opt.(d)
      Gnuplot Plotting File:
      C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Wu2013DONb_Opt.plt
      Mon Oct 17 11:36:24 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{Background} + (1 - \text{Background}) * \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Log}(\text{Dose})),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 5  
 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

User has chosen the log transformed model

Default Initial (and Specified) Parameter Values

background = 0  
 intercept = 2.37622  
 slope = 0.730575

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | 1         | 1     |
| slope     | 1         | 1     |

Parameter Estimates

| Interval    | Variable   | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|----------|-----------|-----------------------|-------|
| Conf. Limit |            |          |           | Lower                 | Upper |
|             | background | 0        | NA        |                       |       |
| 4226.56     | intercept  | 13.5474  | 2149.54   | -4199.47              |       |
| 1410.54     | slope      | 4.19931  | 717.533   | -1402.14              |       |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance     | Test d.f. | P-value |
|---------------|-----------------|-----------|--------------|-----------|---------|
| Full model    | -2.70337        | 5         |              |           |         |
| Fitted model  | -2.70337        | 2         | 4.23246e-008 | 3         | 1       |
| Reduced model | -20.527         | 1         | 35.6472      | 4         | <.0001  |
| AIC:          | 9.40673         |           |              |           |         |

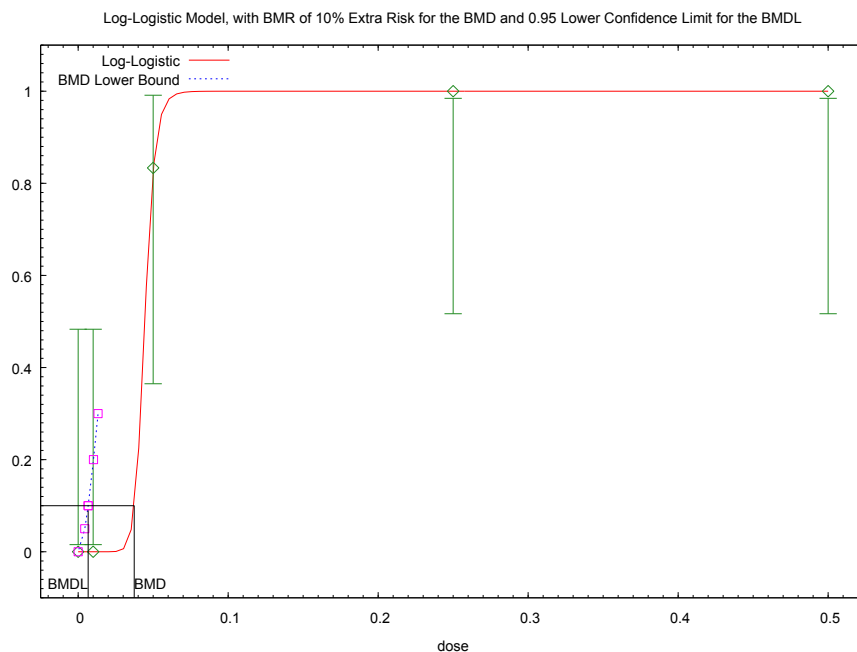
Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|--------|------------|----------|----------|-------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 6.000 | 0.000           |
| 0.0100 | 0.0000     | 0.000    | 0.000    | 6.000 | -0.000          |
| 0.0500 | 0.8333     | 5.000    | 5.000    | 6.000 | 0.000           |
| 0.2500 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |
| 0.5000 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |

Chi<sup>2</sup> = 0.00      d.f. = 3      P-value = 1.0000

Benchmark Dose Computation

Specified effect =                    0.1  
 Risk Type                    =        Extra risk  
 Confidence level =                    0.95  
                                           BMD =            0.0292672  
                                           BMDL =           0.00672434



11:37 10/17 2016

```
=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
      C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Wu2013DONb_Opt.(d)
      Gnuplot Plotting File:
      C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Wu2013DONb_Opt.plt
      Mon Oct 17 11:37:52 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect  
 Independent variable = Dose  
 Slope parameter is not restricted

Total number of observations = 5  
 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



User has chosen the log transformed model

Default Initial Parameter Values

```
background = 0
intercept = 4.15144
slope = 1.28419
```

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix )

|           | intercept | slope |
|-----------|-----------|-------|
| intercept | NA        | NA    |
| slope     | NA        | NA    |

NA - This parameter's variance has been estimated as zero or less.  
THE MODEL HAS PROBABLY NOT CONVERGED!!!

Parameter Estimates

| Interval    | Variable   | Estimate | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|----------|-----------|-----------------------|-------|
|             |            |          |           | Lower Conf. Limit     | Upper |
| Conf. Limit | background | 0        | NA        |                       |       |
| NA          | intercept  | 40.8212  | NA        | NA                    |       |
| NA          | slope      | 13.0892  | NA        | NA                    |       |

At least some variance estimates are negative.  
THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!  
Try again from another starting point.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance     | Test d.f. | P-value |
|---------------|-----------------|-----------|--------------|-----------|---------|
| Full model    | -2.70337        | 5         |              |           |         |
| Fitted model  | -2.70337        | 2         | 4.56654e-008 | 3         | 1       |
| Reduced model | -20.527         | 1         | 35.6472      | 4         | <.0001  |

AIC: 9.40673

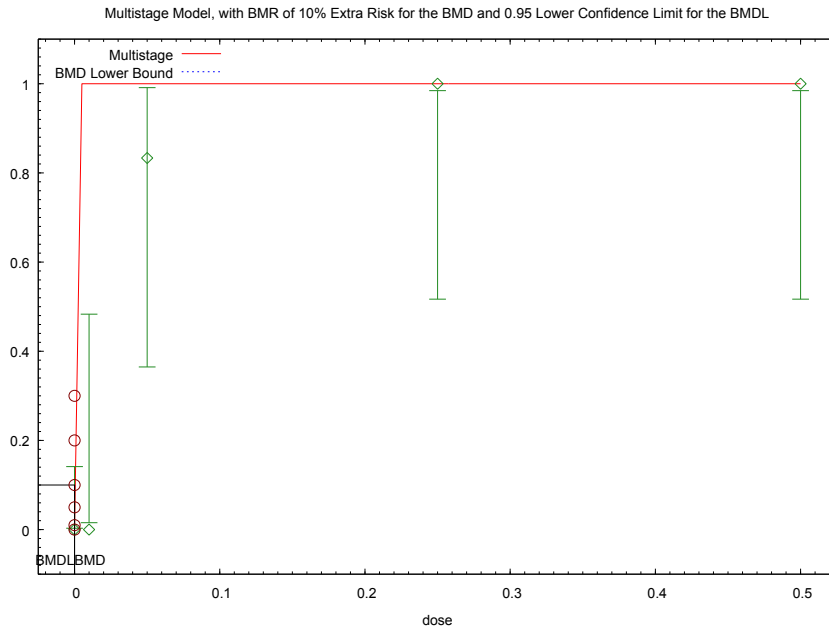
Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size  | Scaled Residual |
|--------|------------|----------|----------|-------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 6.000 | 0.000           |
| 0.0100 | 0.0000     | 0.000    | 0.000    | 6.000 | -0.000          |
| 0.0500 | 0.8333     | 5.000    | 5.000    | 6.000 | 0.000           |
| 0.2500 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |
| 0.5000 | 1.0000     | 6.000    | 6.000    | 6.000 | 0.000           |

Chi^2 = 0.00      d.f. = 3      P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 0.0373825  
 BMDL = 0.00667249



11:39 10/17 2016

```
=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Wu2013DON_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Wu2013DON_Opt.plt
Mon Oct 17 11:39:12 2016
=====
```

BMDS\_Model\_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{beta1} * \text{dose}^1 - \text{beta2} * \text{dose}^2)]$$

The parameter betas are not restricted

Dependent variable = Effect  
 Independent variable = Dose

Total number of observations = 5  
 Total number of records with missing values = 0  
 Total number of parameters in model = 3  
 Total number of specified parameters = 0  
 Degree of polynomial = 2

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

Background = 0  
 Beta(1) = 6.04798e+020  
 Beta(2) = -7.64845e+020

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )

|         | Beta(1) | Beta(2) |
|---------|---------|---------|
| Beta(1) | 1       | 0       |
| Beta(2) | 0       | 1       |

Parameter Estimates

| Interval    | Variable   | Estimate     | Std. Err. | 95.0% Wald Confidence |       |
|-------------|------------|--------------|-----------|-----------------------|-------|
|             |            |              |           | Lower Conf. Limit     | Upper |
| Conf. Limit | Background | 0            | NA        |                       |       |
|             | Beta(1)    | 6.04985e+020 | 1         | 6.04985e+020          |       |
|             | Beta(2)    | -7.6491e+020 | 1         | -7.6491e+020          | -     |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -2.70337        | 5         |          |           |         |
| Fitted model  | -4970           | 2         | 9934.59  | 3         | 0       |
| Reduced model | -33.6365        | 1         | 61.8664  | 4         | <.0001  |

AIC: 9944

Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size   | Scaled Residual |
|--------|------------|----------|----------|--------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 30.000 | -0.008          |
| 0.0100 | 1.0000     | 6.000    | 0.000    | 6.000  | -6.000          |
| 0.0500 | 1.0000     | 6.000    | 5.000    | 6.000  | -1.000          |
| 0.2500 | 1.0000     | 6.000    | 6.000    | 6.000  | 0.000           |
| 0.5000 | 1.0000     | 6.000    | 6.000    | 6.000  | 0.000           |

Chi^2 = 37.00      d.f. = 3      P-value = 0.0000



Total number of specified parameters = 0  
 Degree of polynomial = 2

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

Background = 1  
 Beta(1) = 2.31673e+020  
 Beta(2) = 0

Asymptotic Correlation Matrix of Parameter Estimates

( \*\*\* The model parameter(s) -Background -Beta(1)  
 have been estimated at a boundary point, or have been specified by  
 the user,  
 and do not appear in the correlation matrix )  
 Beta(2)  
 Beta(2) 1

Parameter Estimates

|          |            | 95.0% Wald Confidence |           |                   |                   |
|----------|------------|-----------------------|-----------|-------------------|-------------------|
| Interval | Variable   | Estimate              | Std. Err. | Lower Conf. Limit | Upper Conf. Limit |
|          | Background | 0                     | NA        |                   |                   |
|          | Beta(1)    | 0                     | NA        |                   |                   |
| 1276.56  | Beta(2)    | 646.345               | 321.543   | 16.1312           |                   |

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

| Model         | Log(likelihood) | # Param's | Deviance | Test d.f. | P-value |
|---------------|-----------------|-----------|----------|-----------|---------|
| Full model    | -2.70337        | 5         |          |           |         |
| Fitted model  | -3.11139        | 1         | 0.816043 | 4         | 0.9363  |
| Reduced model | -33.6365        | 1         | 61.8664  | 4         | <.0001  |

AIC: 8.22278

Goodness of Fit

| Dose   | Est._Prob. | Expected | Observed | Size   | Scaled Residual |
|--------|------------|----------|----------|--------|-----------------|
| 0.0000 | 0.0000     | 0.000    | 0.000    | 30.000 | 0.000           |
| 0.0100 | 0.0626     | 0.376    | 0.000    | 6.000  | -0.633          |
| 0.0500 | 0.8013     | 4.808    | 5.000    | 6.000  | 0.197           |
| 0.2500 | 1.0000     | 6.000    | 6.000    | 6.000  | 0.000           |
| 0.5000 | 1.0000     | 6.000    | 6.000    | 6.000  | 0.000           |

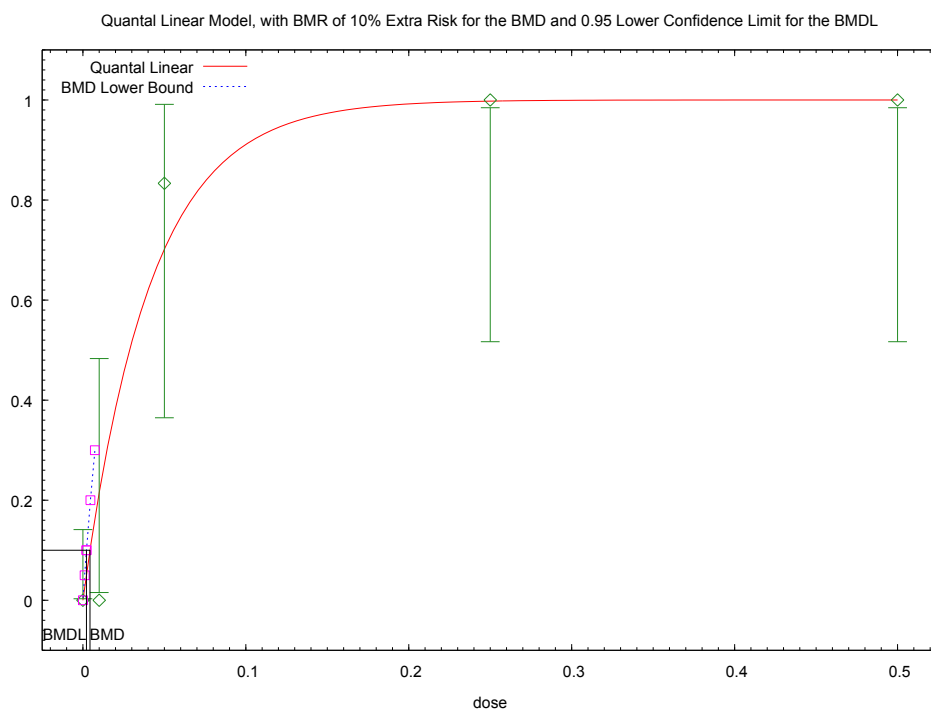
Chi^2 = 0.44      d.f. = 4      P-value = 0.9791

Benchmark Dose Computation

Specified effect = 0.1  
 Risk Type = Extra risk  
 Confidence level = 0.95  
 BMD = 0.0127675  
 BMDL = 0.00369355  
 BMDU = 0.020009

Taken together, (0.00369355, 0.020009) is a 90 % two-sided confidence interval for the BMD

Cancer Slope Factor = 27.0742



11:40 10/17 2016

```

=====
Quantal Linear Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Wu2013DON_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Wu2013DON_Opt.plt
Mon Oct 17 11:40:55 2016
=====
    
```

BMDS\_Model\_Run  
 ~~~~~

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose})]$$

Dependent variable = Effect
 Independent variable = Dose

Total number of observations = 5
 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 (and Specified) Parameter Values
 Background = 0.03125
 Slope = 4.09539
 Power = 1 Specified

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Power
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)
 Slope
 Slope 1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
45.4486	Slope	24.1749	10.8541	2.90119	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-4.44671	1	3.48668	4	0.4799
Reduced model	-33.6365	1	61.8664	4	<.0001

AIC: 10.8934

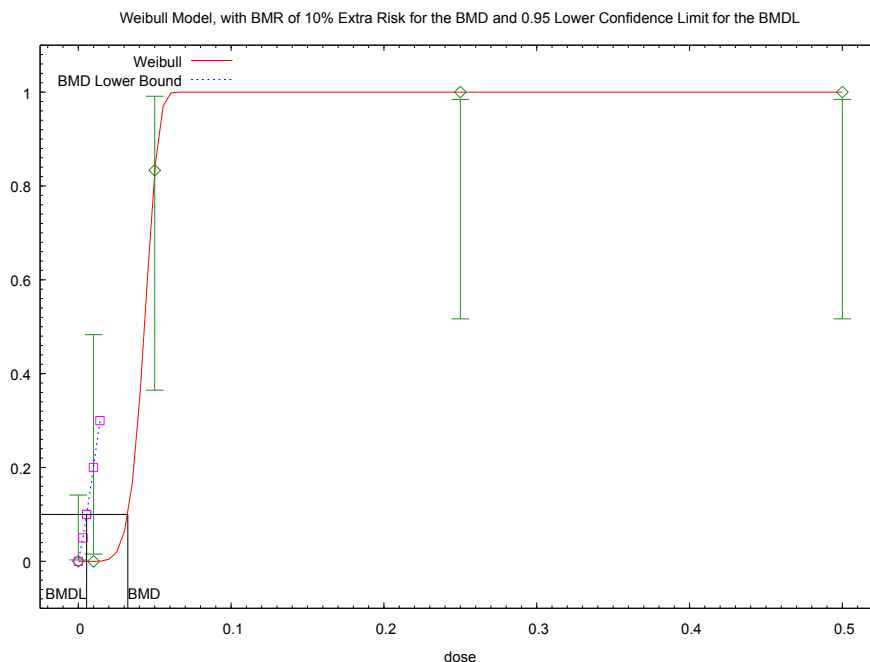
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	30.000	0.000
0.0100	0.2147	1.288	0.000	6.000	-1.281
0.0500	0.7014	4.209	5.000	6.000	0.706
0.2500	0.9976	5.986	6.000	6.000	0.119
0.5000	1.0000	6.000	6.000	6.000	0.006

Chi^2 = 2.15 d.f. = 4 P-value = 0.7075

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.00435826
 BMDL = 0.00217173



11:41 10/17 2016

```
=====
Weibull Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Wu2013DON_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Wu2013DON_Opt.plt
Mon Oct 17 11:41:52 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose}^{\text{power}})]$$

Dependent variable = Effect
 Independent variable = Dose
 Power parameter is not restricted

Total number of observations = 5
 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 (and Specified) Parameter Values

Background = 0.03125
 Slope = 2.81431
 Power = 0.458784

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	Slope	Power
Slope	1	1
Power	1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper Conf. Limit
	Background	0	NA		
7.72159e+010	Slope	5.71235e+008	3.91051e+010	-7.60734e+010	
51.3227	Power	6.53604	22.8508	-38.2506	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.70366	2	0.000580701	3	1
Reduced model	-33.6365	1	61.8664	4	<.0001
AIC:	9.40732				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	30.000	0.000
0.0100	0.0000	0.000	0.000	6.000	-0.017
0.0500	0.8333	5.000	5.000	6.000	0.000
0.2500	1.0000	6.000	6.000	6.000	0.000
0.5000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.00 d.f. = 3 P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95

BMD = 0.0324114

BMDL = 0.00546056

Supplementary information for WUDON3AC (Wu et al., 2013a)

Details of the BMD Analysis of concentration-response data for vomiting in mink of Wu et al. (2013) reported in Appendix G, Section G.4.3.2 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed - for 3-Ac-DON

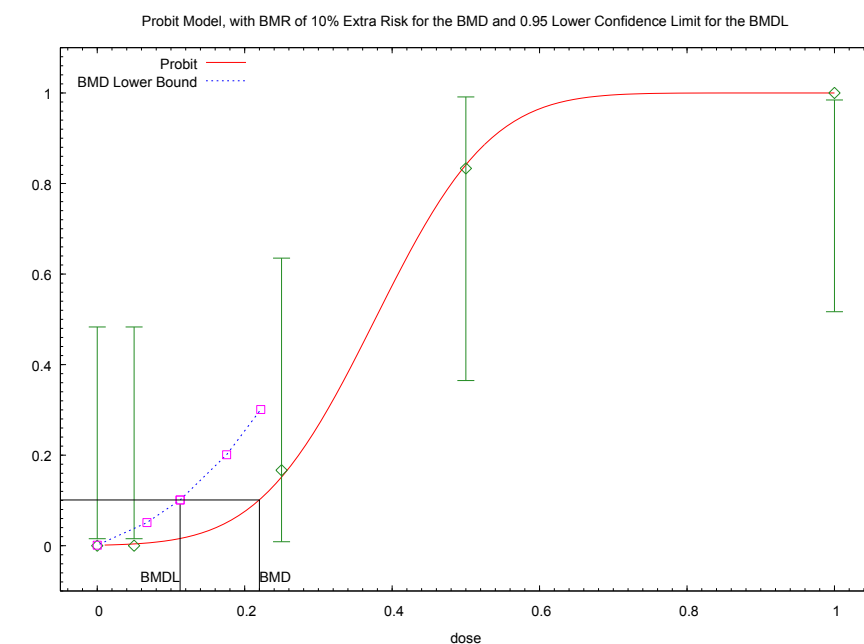
Filename: WuDON3Ac

Folder: Mink

Output from BMDS from 19 Oct 2016 with the full information of the fit of all models of BMDS where a model fit was obtained in a reasonable duration of computing time.

The output is ordered in the sequence used in the Table G18 shown in Appendix G which is based on an order of increasing mathematical complexity. The two less complex multistage models are nested under the multistage model which was used with the default assumption of the degree of the polynomial of BMDS.

Occasionally, it was not possible to obtain the results of the Gamma model with the notebook used for the calculation HP844op or Lenovo 3508. This was the case for the data of 3-Ac-DON. The Gamma model failed without any output. The Multistage model run but was considered to failed since a statistically impossible p-value of 2 was recorded.



```
=====
Probit Model. (Version: 3.3; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Wu2013-3AcDONb_Opt. (d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/pro_Wu2013-3AcDONb_Opt.plt
Mon Oct 17 17:29:49 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Dose}),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 5
 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 (and Specified) Parameter Values
 background = 0 Specified
 intercept = -1.82112
 slope = 3.90837

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-0.93
slope	-0.93	1

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence	
				Lower Conf. Limit	Upper
Conf. Limit	intercept	-3.0611	1.16433	-5.34315	-
0.779053	slope	8.11591	3.00731	2.22169	
14.0101					

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-5.44392	2	0.0743692	3	0.9947
Reduced model	-20.1904	1	29.5672	4	<.0001
AIC:	14.8878				

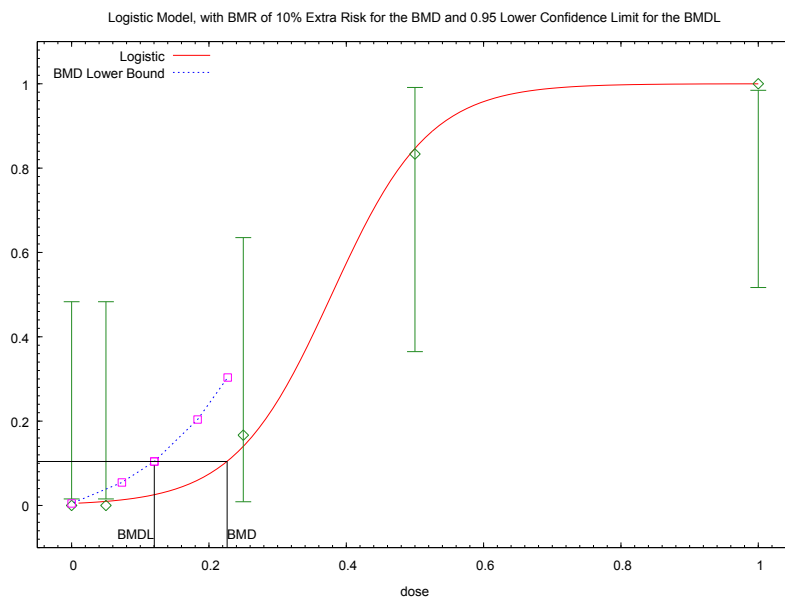
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0011	0.007	0.000	6.000	-0.081
0.0500	0.0040	0.024	0.000	6.000	-0.154
0.2500	0.1510	0.906	1.000	6.000	0.107
0.5000	0.8406	5.043	5.000	6.000	-0.049
1.0000	1.0000	6.000	6.000	6.000	0.001

Chi² = 0.04 d.f. = 3 P-value = 0.9976

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.219961
 BMDL = 0.112409



17:34 10/17 2016

```
=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
      C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Wu2013-3AcDONb_Opt. (d)
      Gnuplot Plotting File:
      C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Wu2013-3AcDONb_Opt.plt
      Mon Oct 17 17:34:55 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = 1/[1+\text{EXP}(-\text{intercept}-\text{slope}*\text{dose})]$$

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 5
 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
background = 0 Specified
intercept = -2.50184
slope = 5.52459

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-0.93
slope	-0.93	1

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence	
				Lower Conf. Limit	Upper
Conf. Limit	intercept	-5.34289	2.12811	-9.51391	-
1.17188	slope	14.0965	5.45861	3.39782	
24.7952					

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-5.51526	2	0.217047	3	0.9748
Reduced model	-20.1904	1	29.5672	4	<.0001
AIC:	15.0305				

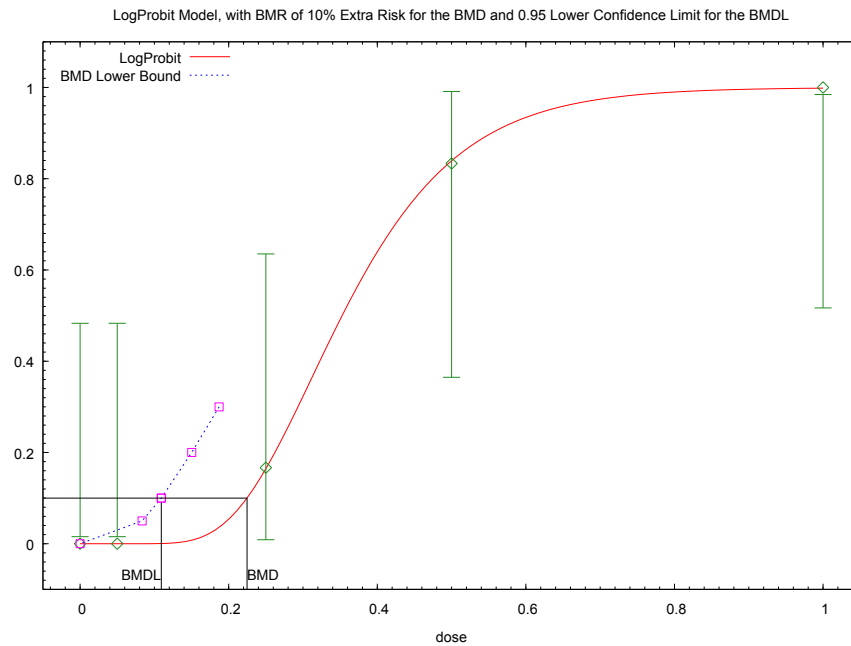
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0048	0.029	0.000	6.000	-0.169
0.0500	0.0096	0.058	0.000	6.000	-0.241
0.2500	0.1396	0.837	1.000	6.000	0.191
0.5000	0.8462	5.077	5.000	6.000	-0.088
1.0000	0.9998	5.999	6.000	6.000	0.031

Chi^2 = 0.13 d.f. = 3 P-value = 0.9877

Benchmark Dose Computation

Specified effect = 0.1
Risk Type = Extra risk
Confidence level = 0.95
BMD = 0.226466
BMDL = 0.120421



```

=====
      Probit Model. (Version: 3.3; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Wu2013-3AcDONb_Opt. (d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Wu2013-3AcDONb_Opt.plt
      Mon Oct 17 17:36:45 2016
=====
  
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{Background} + (1 - \text{Background}) * \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Log}(\text{Dose})),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 5
 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

User has chosen the log transformed model

```

Default Initial (and Specified) Parameter Values
background = 0
intercept = 1.30032
slope = 1.02485
  
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	intercept	slope
intercept	1	0.94
slope	0.94	1

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence	
Conf. Limit				Lower	Upper
	background	0	NA		
5.44123	intercept	2.95845	1.26675	0.475665	
5.12227	slope	2.84	1.16444	0.557738	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-5.41687	2	0.0202628	3	0.9992
Reduced model	-20.1904	1	29.5672	4	<.0001
AIC:	14.8337				

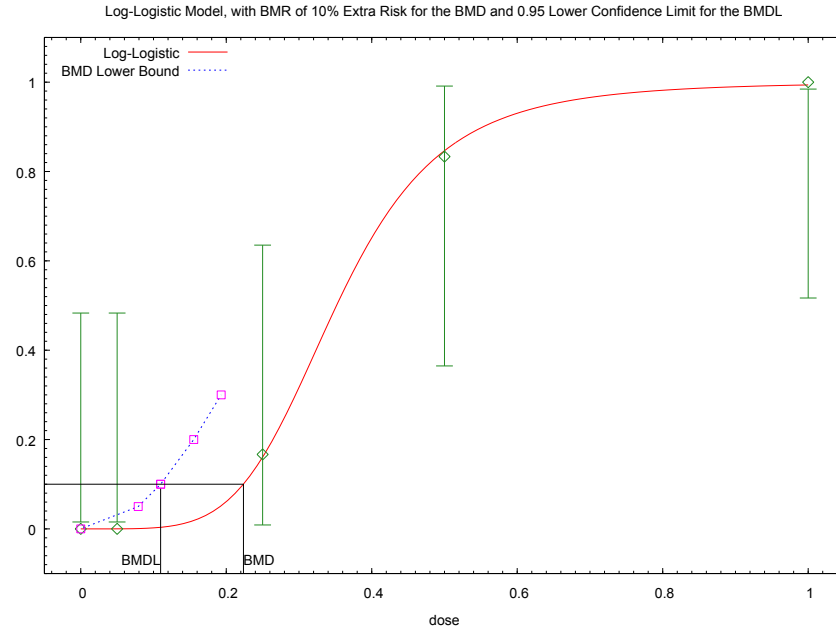
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0500	0.0000	0.000	0.000	6.000	-0.000
0.2500	0.1639	0.983	1.000	6.000	0.018
0.5000	0.8389	5.033	5.000	6.000	-0.037
1.0000	0.9985	5.991	6.000	6.000	0.096

Chi^2 = 0.01 d.f. = 3 P-value = 0.9997

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.224708
 BMDL = 0.109274



17:38 10/17 2016

```

=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Wu2013-3AcDONb_Opt. (d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Wu2013-3AcDONb_Opt.plt
                                          Mon Oct 17 17:38:39 2016
=====
  
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 5
 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

User has chosen the log transformed model

```

      Default Initial Parameter Values
      background =          0
      intercept =    2.26129
      slope =          1.78223
  
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background

the user, have been estimated at a boundary point, or have been specified by
and do not appear in the correlation matrix)

	intercept	slope
intercept	1	0.94
slope	0.94	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	background	0	NA		
9.49414	intercept	5.06531	2.25965	0.636483	
8.91426	slope	4.8493	2.074	0.784331	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-5.44955	2	0.0856239	3	0.9935
Reduced model	-20.1904	1	29.5672	4	<.0001
AIC:	14.8991				

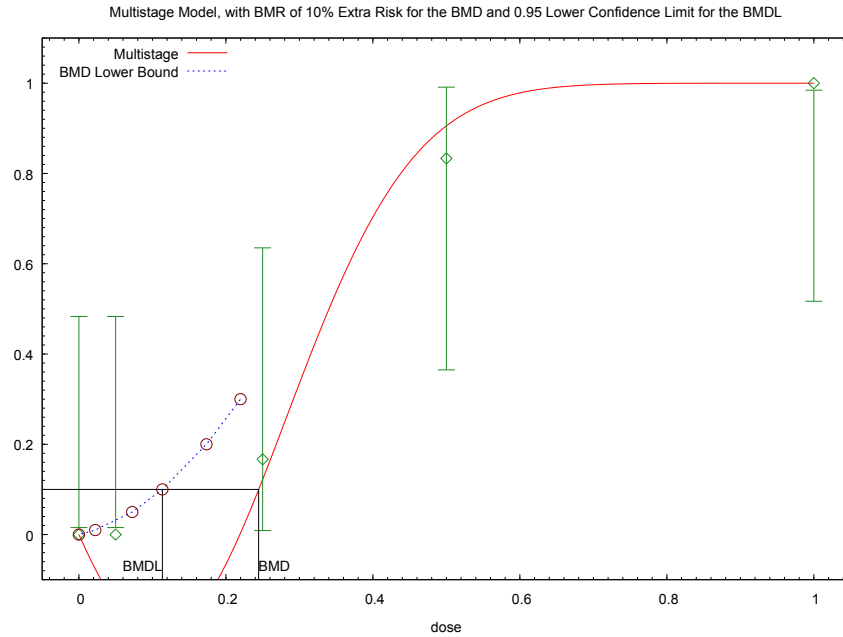
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0500	0.0001	0.000	0.000	6.000	-0.022
0.2500	0.1601	0.961	1.000	6.000	0.044
0.5000	0.8461	5.076	5.000	6.000	-0.086
1.0000	0.9937	5.962	6.000	6.000	0.195

Chi^2 = 0.05 d.f. = 3 P-value = 0.9973

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.223656
 BMDL = 0.109804



```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Wu2013-3AcDONb_Opt. (d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Wu2013-3AcDONb_Opt.plt
Mon Oct 17 17:40:08 2016
=====

```

```

BMDS_Model_Run
~~~~~

```

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{beta1} * \text{dose}^{\text{beta2}})]$$

The parameter betas are not restricted

Dependent variable = Effect
 Independent variable = Dose

```

Total number of observations = 5
Total number of records with missing values = 0
Total number of parameters in model = 3
Total number of specified parameters = 0
Degree of polynomial = 2

```

```

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
Background = 1
Beta(1) = -8.69869e+019
Beta(2) = 1.82843e+020

```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	Beta(1)	Beta(2)
Beta(1)	1	-0.98
Beta(2)	-0.98	1

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence Lower Conf. Limit	Upper
	Background	0	NA		
1.84608	Beta(1)	-3.679	2.81897	-9.20408	
37.7586	Beta(2)	16.8077	10.6894	-4.1432	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Warning: Likelihood for the fitted model larger than the Likelihood for the full model.

Error in computing chi-square; returning 2

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-4.75841	2	-1.29665	3	2
Reduced model	-20.1904	1	29.5672	4	<.0001
AIC:	13.5168				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0500	-0.1525	-0.915	0.000	6.000	0.915
0.2500	0.1225	0.735	1.000	6.000	0.330
0.5000	0.9058	5.435	5.000	6.000	-0.608
1.0000	1.0000	6.000	6.000	6.000	0.003

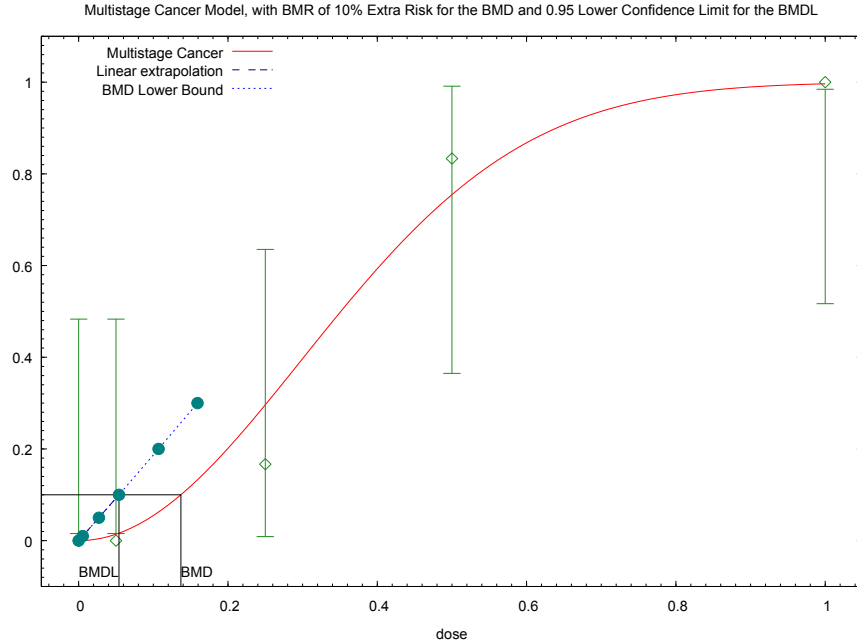
Chi^2 = 1.32 d.f. = 3 P-value = 0.7255

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.244523

BMDL = 0.113603
 BMDU = 0.27287

Taken together, (0.113603, 0.27287) is a 90 % two-sided confidence interval for the BMD



17:42 10/17 2016

```
=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Wu2013-3AcDONb_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Wu2013-3AcDONb_Opt.plt
Mon Oct 17 17:42:48 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1 - \text{EXP}(-\text{beta1} * \text{dose}^{1-\text{beta2}} * \text{dose}^2)]$$

The parameter betas are restricted to be positive

Dependent variable = Effect
 Independent variable = Dose

Total number of observations = 5
 Total number of records with missing values = 0
 Total number of parameters in model = 3
 Total number of specified parameters = 0
 Degree of polynomial = 2

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

Background = 0
 Beta(1) = 0
 Beta(2) = 1.0228e+020

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Beta(1) have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

Beta(2)
 Beta(2) 1

Parameter Estimates

95.0% Wald Confidence

Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
	Beta(1)	0	NA		
10.1336	Beta(2)	5.61613	2.30489	1.09863	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-5.8917	1	0.969934	4	0.9143
Reduced model	-20.1904	1	29.5672	4	<.0001
AIC:	13.7834				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0500	0.0139	0.084	0.000	6.000	-0.291
0.2500	0.2960	1.776	1.000	6.000	-0.694
0.5000	0.7544	4.526	5.000	6.000	0.449
1.0000	0.9964	5.978	6.000	6.000	0.148

Chi^2 = 0.79 d.f. = 4 P-value = 0.9397

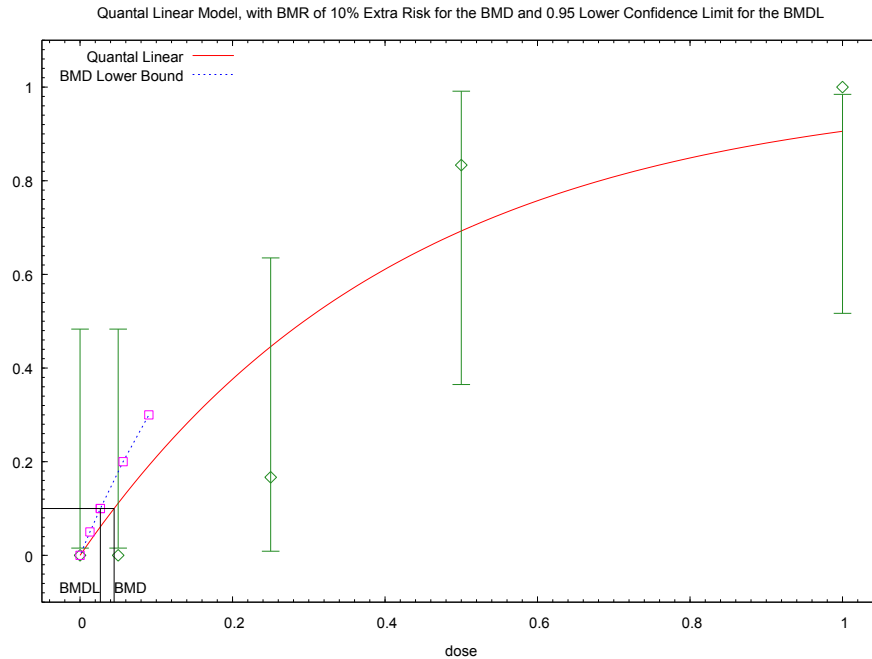
Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.136968

BMDL = 0.0540174
 BMDU = 0.193732

Taken together, (0.0540174, 0.193732) is a 90 % two-sided confidence interval for the BMD

Cancer Slope Factor = 1.85126



17:45 10/17 2016

```
=====
Quantal Linear Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Wu2013-3AcDONb_Opt. (d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Wu2013-3AcDONb_Opt.plt
Mon Oct 17 17:45:28 2016
=====
```

```
BMDS_Model_Run
~~~~~
```

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) * [1 - \text{EXP}(-\text{slope} * \text{dose})]$$

Dependent variable = Effect
 Independent variable = Dose

Total number of observations = 5
 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 (and Specified) Parameter Values
 Background = 0.125

Slope = 1.94591
 Power = 1 Specified

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Power
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

Slope
 Slope 1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
	Slope	2.3573	0.773732	0.840811	
				3.87379	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-8.07688	1	5.34029	4	0.2541
Reduced model	-20.1904	1	29.5672	4	<.0001
AIC:	18.1538				

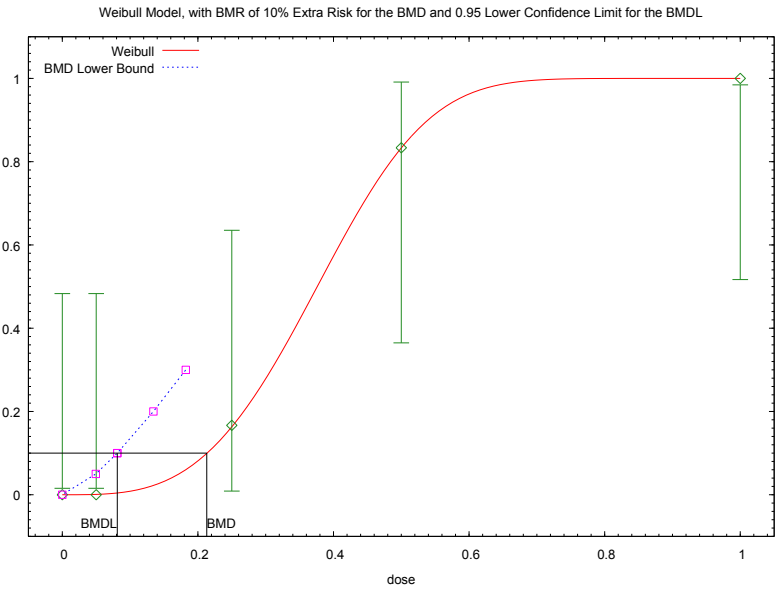
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0500	0.1112	0.667	0.000	6.000	-0.866
0.2500	0.4453	2.672	1.000	6.000	-1.373
0.5000	0.6923	4.154	5.000	6.000	0.748
1.0000	0.9053	5.432	6.000	6.000	0.792

Chi^2 = 3.82 d.f. = 4 P-value = 0.4303

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.0446954
 BMDL = 0.026661



```

=====
      Weibull Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Wu2013-3AcDONb_Opt. (d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Wu2013-3AcDONb_Opt.plt
      Mon Oct 17 17:47:01 2016
=====
  
```

```

BMDS_Model_Run
~~~~~
  
```

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose}^{\text{power}})]$$

Dependent variable = Effect
 Independent variable = Dose
 Power parameter is not restricted

Total number of observations = 5
 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 (and Specified) Parameter Values
      Background =      0.125
      Slope =         1.94591
      Power =         1.27059
  
```

```

      Asymptotic Correlation Matrix of Parameter Estimates

      ( *** The model parameter(s) -Background
            have been estimated at a boundary point, or have been specified by
the user,
            and do not appear in the correlation matrix )

      Slope      Power
  
```

Slope 1 0.95
 Power 0.95 1

Parameter Estimates

95.0% Wald Confidence

Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
67.4214	Slope	18.0225	25.204	-31.3764	
6.42723	Power	3.32593	1.58233	0.224626	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-5.40673	5			
Fitted model	-5.41199	2	0.0105094	3	0.9997
Reduced model	-20.1904	1	29.5672	4	<.0001

AIC: 14.824

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0500	0.0008	0.005	0.000	6.000	-0.071
0.2500	0.1641	0.985	1.000	6.000	0.017
0.5000	0.8342	5.005	5.000	6.000	-0.006
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.01 d.f. = 3 P-value = 0.9999

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.213092
 BMDL = 0.0810851

Gamma Model failed

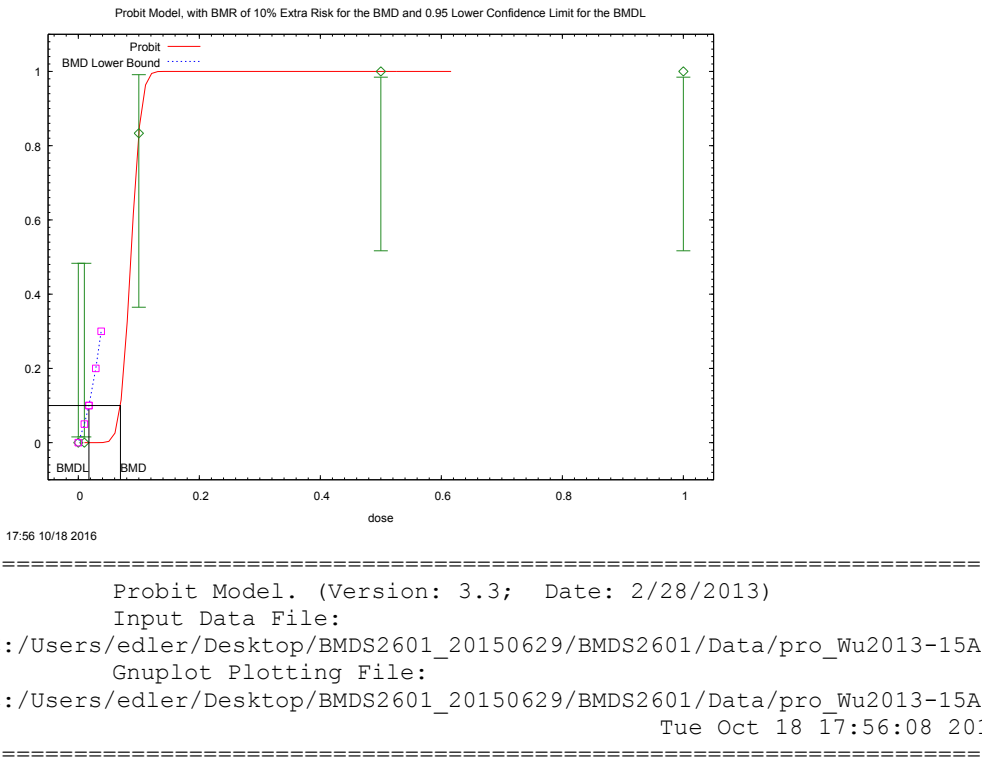
Supplementary information for WUDON15AC (Wu et al., 2013a)

Details of the BMD Analysis of concentration-response data for vomiting in mink of Wu et al. (2013) reported in Appendix G, Section G.4.3.3 of the Scientific Opinion on the risks to human and animal health related to the presence of deoxynivalenol and its acetylated and modified forms in food and feed - for 15-Ac-DON

Filename:WuDON15Ac Folder: Mink

Output from BMDS from 19 Oct 2016 with the full information of the fit of all models of BMDS where a model fit was obtained in a reasonable duration of computing time.

The output is ordered in the sequence used in the Table G19 shown in Appendix G which is based on an order of increasing mathematical complexity. The two less complex multistage models are nested under the multistage model which was used with the default assumption of the degree of the polynomial of BMDS.



```

BMDS_Model_Run
~~~~~

The form of the probability function is:

P[response] = CumNorm(Intercept+Slope*Dose),

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect
Independent variable = Dose
Slope parameter is not restricted

Total number of observations = 5
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 (and Specified) Parameter Values
 background = 0 Specified
 intercept = -0.970097
 slope = 3.37034

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

	intercept	slope
intercept	1	-1
slope	-1	1

Parameter Estimates

Interval	Variable	Estimate	Std. Err.	95.0% Wald Confidence	
				Lower Conf. Limit	Upper
2468.34	intercept	-6.4392	1262.67	-2481.22	
24821.9	slope	74.0661	12626.7	-24673.8	

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.70337	2	7.39017e-008	3	1
Reduced model	-20.527	1	35.6472	4	<.0001
AIC:	9.40673				

Goodness of Fit

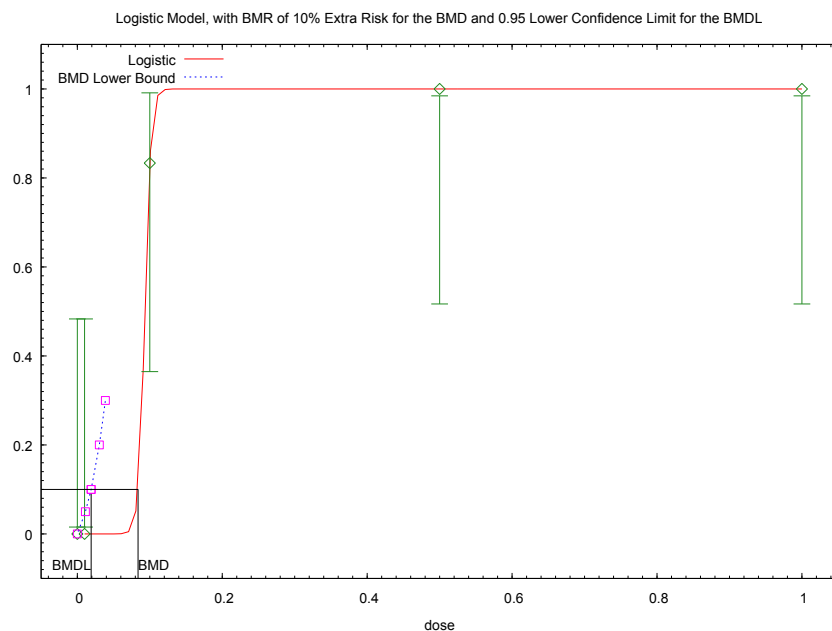
Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	-0.000
0.0100	0.0000	0.000	0.000	6.000	-0.000
0.1000	0.8333	5.000	5.000	6.000	0.000
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.00 d.f. = 3 P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk

Confidence level = 0.95
 BMD = 0.0696358
 BMDL = 0.0175236



18:00 10/18 2016

```
=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Wu2013-15AcDONb_Opt.(d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/log_Wu2013-15AcDONb_Opt.plt
      Tue Oct 18 18:00:05 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = 1/[1+\text{EXP}(-\text{intercept}-\text{slope}*\text{dose})]$$

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 5
 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
background = 0 Specified
intercept = -1.27414
slope = 4.76396
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background
 have been estimated at a boundary point, or have been specified by
 the user,
 and do not appear in the correlation matrix)

	intercept	slope
intercept	NA	NA
slope	NA	NA

NA - This parameter's variance has been estimated as zero or less.
 THE MODEL HAS PROBABLY NOT CONVERGED!!!

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	intercept	-21.8671	NA	NA	NA
NA	slope	234.766	NA	NA	NA
NA					

At least some variance estimates are negative.
 THIS USUALLY MEANS THE MODEL HAS NOT CONVERGED!
 Try again from another starting point.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.70337	2	4.47561e-008	3	1
Reduced model	-20.527	1	35.6472	4	<.0001
AIC:	9.40673				

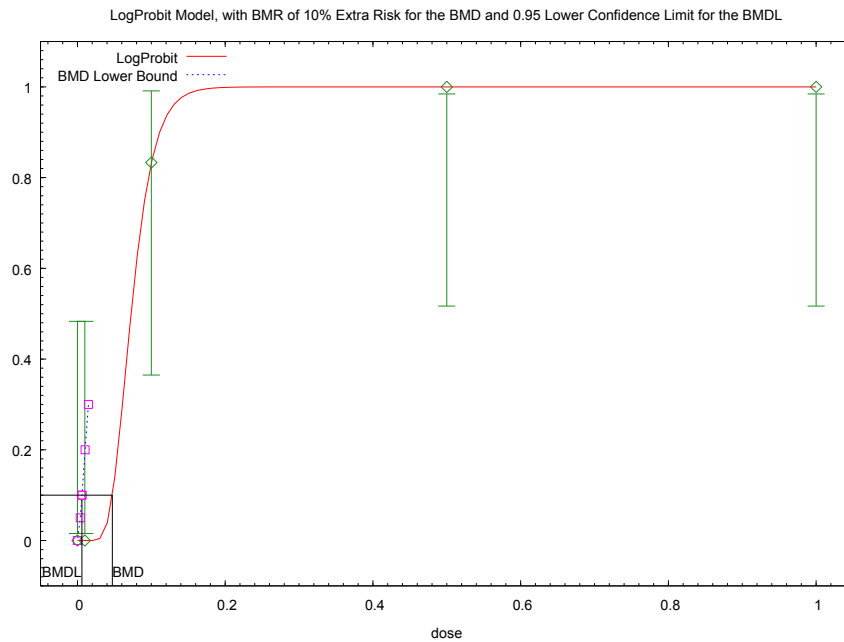
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	-0.000
0.0100	0.0000	0.000	0.000	6.000	-0.000
0.1000	0.8333	5.000	5.000	6.000	-0.000
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.00 d.f. = 3 P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.0837851
 BMDL = 0.0190577



18:01 10/18 2016

```

=====
      Probit Model. (Version: 3.3; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Wu2013-15AcDONb_Opt. (d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnp_Wu2013-15AcDONb_Opt.plt
                                          Tue Oct 18 18:01:37 2016
=====

```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{Background} + (1 - \text{Background}) * \text{CumNorm}(\text{Intercept} + \text{Slope} * \text{Log}(\text{Dose})),$$

where CumNorm(.) is the cumulative normal distribution function

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 5
 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

User has chosen the log transformed model

```

Default Initial (and Specified) Parameter Values
background = 0
intercept = 1.83904
slope = 0.647754

```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

	intercept	slope
intercept	1	1
slope	1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	background	0	NA		
3229.89	intercept	7.87447	1643.91	-3214.14	
1402.3	slope	2.9997	713.943	-1396.3	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.70337	2	5.81484e-008	3	1
Reduced model	-20.527	1	35.6472	4	<.0001
AIC:	9.40673				

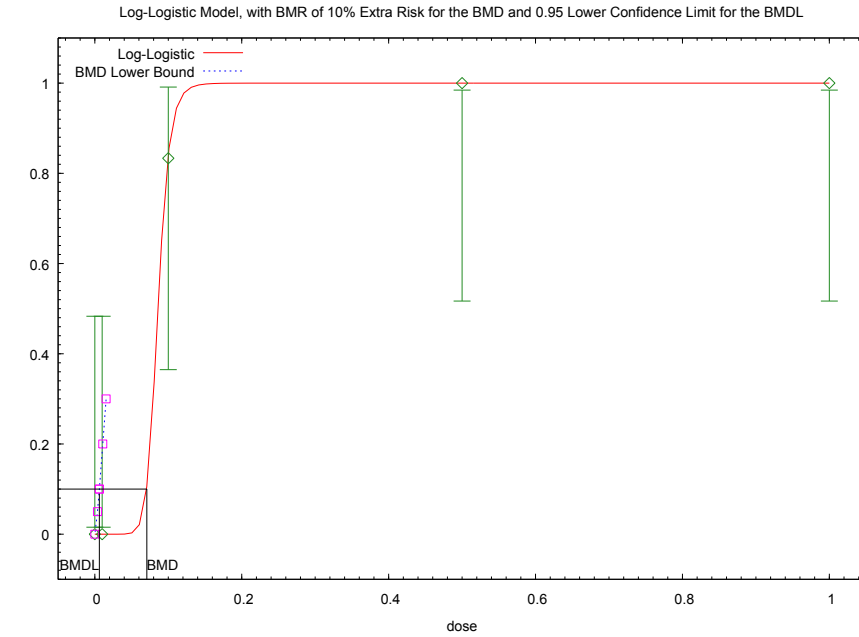
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0100	0.0000	0.000	0.000	6.000	-0.000
0.1000	0.8333	5.000	5.000	6.000	0.000
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.00 d.f. = 3 P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.0472493
 BMDL = 0.00603745



```

=====
      Logistic Model. (Version: 2.14; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Wu2013-15AcDONb_Opt. (d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/lnl_Wu2013-15AcDONb_Opt.plt
                                          Tue Oct 18 18:03:16 2016
=====
    
```

```

BMDS_Model_Run
~~~~~
    
```

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1 - \text{background}) / [1 + \text{EXP}(-\text{intercept} - \text{slope} * \text{Log}(\text{dose}))]$$

Dependent variable = Effect
 Independent variable = Dose
 Slope parameter is not restricted

Total number of observations = 5
 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

User has chosen the log transformed model

```

      Default Initial Parameter Values
      background =          0
      intercept  =      3.20342
      slope      =      1.13661
    
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -background

the user, have been estimated at a boundary point, or have been specified by
and do not appear in the correlation matrix)

	intercept	slope
intercept	1	1
slope	1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	background	0	NA		
16555.5	intercept	27.0242	8433.06	-16501.5	
7189.27	slope	11.0375	3662.43	-7167.2	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.70337	2	4.80546e-008	3	1
Reduced model	-20.527	1	35.6472	4	<.0001
AIC:	9.40673				

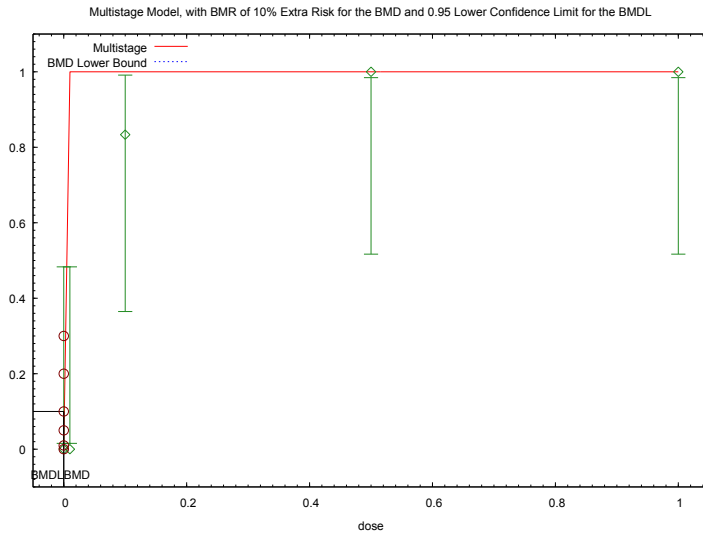
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0100	0.0000	0.000	0.000	6.000	-0.000
0.1000	0.8333	5.000	5.000	6.000	0.000
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.00 d.f. = 3 P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.0708305
 BMDL = 0.00614071



18:04 10/18 2016

```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Wu2013-15AcDONb_Opt.(d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/mst_Wu2013-15AcDONb_Opt.plt
Tue Oct 18 18:04:46 2016
=====

```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1 - \text{EXP}(-\text{beta1} * \text{dose}^1 - \text{beta2} * \text{dose}^2)]$$

The parameter betas are not restricted

Dependent variable = Effect
Independent variable = Dose

Total number of observations = 5
Total number of records with missing values = 0
Total number of parameters in model = 3
Total number of specified parameters = 0
Degree of polynomial = 2

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

Background = 0
Beta(1) = 2.9711e+020
Beta(2) = -1.87103e+020

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background have been estimated at a boundary point, or have been specified by the user,

and do not appear in the correlation matrix)

	Beta(1)	Beta(2)
Beta(1)	1	0
Beta(2)	0	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
	Beta(1)	2.97202e+020	1	2.97202e+020	
2.97202e+020	Beta(2)	-1.87119e+020	1	-1.87119e+020	-
1.87119e+020					

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-4970	2	9934.59	3	0
Reduced model	-20.527	1	35.6472	4	<.0001
AIC:	9944				

Goodness of Fit

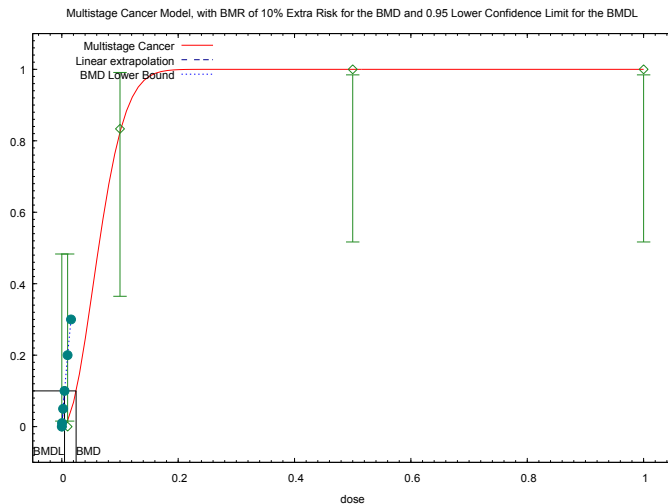
Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	-0.003
0.0100	1.0000	6.000	0.000	6.000	-6.000
0.1000	1.0000	6.000	5.000	6.000	-1.000
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 37.00 d.f. = 3 P-value = 0.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 3.54508e-022
 BMDL = 3.54508e-022
 BMDU = 3.54508e-022

Taken together, (3.54508e-022, 3.54508e-022) is a 90 % two-sided confidence interval for the BMD



18:05 10/18 2016

```

=====
Multistage Model. (Version: 3.4; Date: 05/02/2014)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Wu2013-15AcDONb_Opt. (d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/msc_Wu2013-15AcDONb_Opt.plt
Tue Oct 18 18:05:55 2016
=====
    
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{beta1} * \text{dose}^1 - \text{beta2} * \text{dose}^2)]$$

The parameter betas are restricted to be positive

Dependent variable = Effect
Independent variable = Dose

Total number of observations = 5
Total number of records with missing values = 0
Total number of parameters in model = 3
Total number of specified parameters = 0
Degree of polynomial = 2

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

Background = 1
Beta(1) = 1.15414e+020
Beta(2) = 0

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background -Beta(1) have been estimated at a boundary point, or have been specified by the user, and do not appear in the correlation matrix)

Beta(2)
Beta(2) 1

Parameter Estimates

95.0% Wald Confidence

Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper Conf. Limit
	Background	0	NA		
	Beta(1)	0	NA		
	Beta(2)	174.344	88.2257	1.42469	347.263

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.80941	1	0.212079	4	0.9948
Reduced model	-20.527	1	35.6472	4	<.0001
AIC:	7.61881				

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0100	0.0173	0.104	0.000	6.000	-0.325
0.1000	0.8251	4.950	5.000	6.000	0.053
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

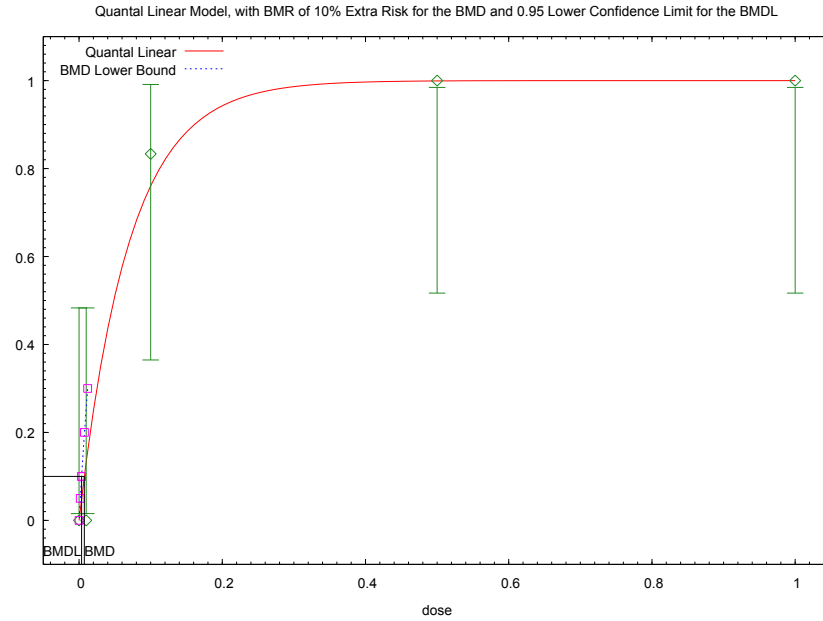
Chi^2 = 0.11 d.f. = 4 P-value = 0.9986

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.024583
 BMDL = 0.004672
 BMDU = 0.0387137

Taken together, (0.004672, 0.0387137) is a 90 % two-sided confidence interval for the BMD

Cancer Slope Factor = 21.4041



18:07 10/18 2016

```
=====
Quantal Linear Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Wu2013-15AcDONb_Opt. (d)
Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/qln_Wu2013-15AcDONb_Opt.plt
Tue Oct 18 18:07:27 2016
=====
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose})]$$

Dependent variable = Effect
Independent variable = Dose

Total number of observations = 5
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 (and Specified) Parameter Values
Background = 0.125
Slope = 1.94591
Power = 1 Specified
```

Asymptotic Correlation Matrix of Parameter Estimates

```
( *** The model parameter(s) -Background -Power
have been estimated at a boundary point, or have been specified by
the user,
and do not appear in the correlation matrix )

Slope
Slope 1
```

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
27.5215	Slope	14.2843	6.7538	1.04714	

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-3.66035	1	1.91397	4	0.7516
Reduced model	-20.527	1	35.6472	4	<.0001

AIC: 9.3207

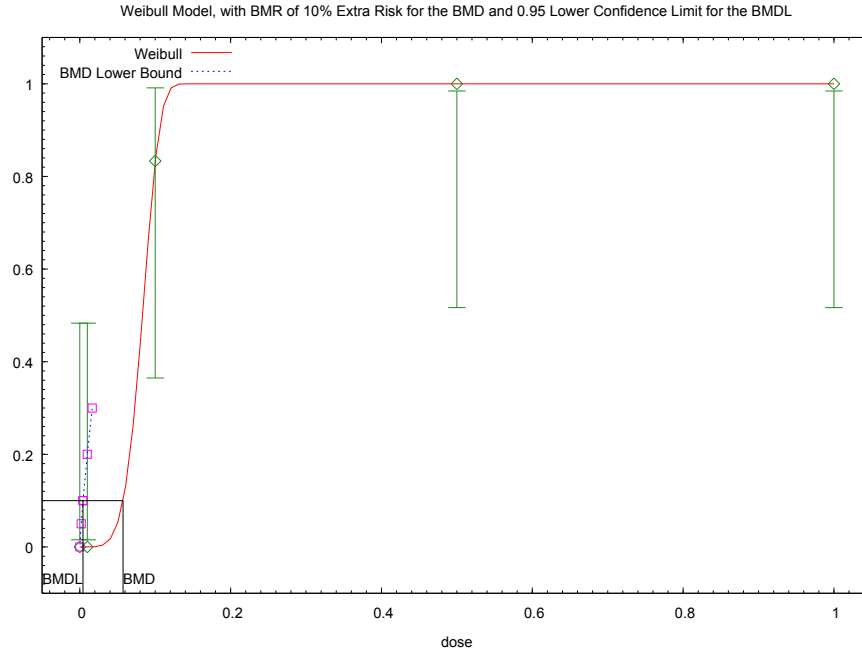
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0100	0.1331	0.799	0.000	6.000	-0.960
0.1000	0.7603	4.562	5.000	6.000	0.419
0.5000	0.9992	5.995	6.000	6.000	0.069
1.0000	1.0000	6.000	6.000	6.000	0.002

Chi² = 1.10 d.f. = 4 P-value = 0.8940

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.00737595
 BMDL = 0.00354948



18:09 10/18 2016

```

=====
      Weibull Model using Weibull Model (Version: 2.16; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Wu2013-15AcDONb_Opt. (d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/wei_Wu2013-15AcDONb_Opt.plt
      Tue Oct 18 18:09:12 2016
=====
    
```

BMDS_Model_Run

The form of the probability function is:

$$P[\text{response}] = \text{background} + (1-\text{background}) * [1-\text{EXP}(-\text{slope} * \text{dose}^{\text{power}})]$$

Dependent variable = Effect
 Independent variable = Dose
 Power parameter is not restricted

Total number of observations = 5
 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 (and Specified) Parameter Values

Background =	0.125
Slope =	1.94591
Power =	0.453362

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background
 have been estimated at a boundary point, or have been specified by
 the user,

and do not appear in the correlation matrix)

	Slope	Power
Slope	1	1
Power	1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper
Conf. Limit	Background	0	NA		
	Slope	224220	1.69921e+007	-3.30798e+007	
3.35282e+007	Power	5.09741	32.9116	-59.4081	
69.6029					

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.70345	2	0.00017181	3	1
Reduced model	-20.527	1	35.6472	4	<.0001
AIC:	9.40691				

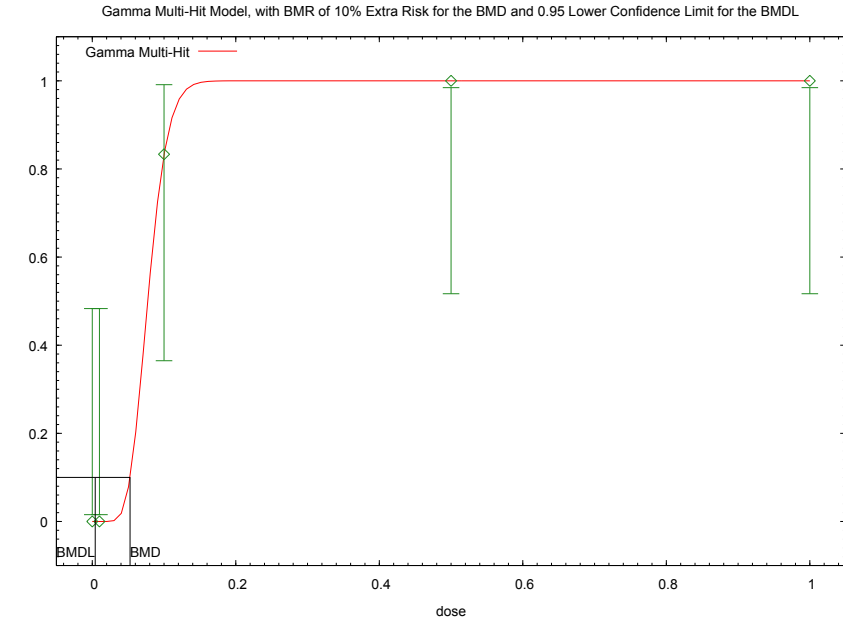
Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0100	0.0000	0.000	0.000	6.000	-0.009
0.1000	0.8333	5.000	5.000	6.000	0.000
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.00 d.f. = 3 P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.0573571
 BMDL = 0.00424741



```

=====
      Gamma Model. (Version: 2.16; Date: 2/28/2013)
      Input Data File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/gam_Wu2013-15AcDONb_Opt. (d)
      Gnuplot Plotting File:
C:/Users/edler/Desktop/BMDS2601_20150629/BMDS2601/Data/gam_Wu2013-15AcDONb_Opt.plt
                                          Tue Oct 18 18:11:14 2016
=====
  
```

```

BMDS_Model_Run
~~~~~
  
```

The form of the probability function is:

$P[\text{response}] = \text{background} + (1 - \text{background}) * \text{CumGamma}[\text{slope} * \text{dose}, \text{power}]$,
 where CumGamma(.) is the cumulative Gamma distribution function

Dependent variable = Effect
 Independent variable = Dose
 Power parameter is not restricted

Total number of observations = 5
 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 (and Specified) Parameter Values
      Background =      0.125
      Slope =          12.8562
      Power =           1.3
  
```

Asymptotic Correlation Matrix of Parameter Estimates

(*** The model parameter(s) -Background
 have been estimated at a boundary point, or have been specified by
 the user, and do not appear in the correlation matrix)

	Slope	Power
Slope	1	1
Power	1	1

Parameter Estimates

		95.0% Wald Confidence			
Interval	Variable	Estimate	Std. Err.	Lower Conf. Limit	Upper Conf. Limit
	Background	0	NA		
	Slope	164.278	19326.5	-37715	38043.5
	Power	13.0048	1701.5	-3321.87	3347.88

NA - Indicates that this parameter has hit a bound implied by some inequality constraint and thus has no standard error.

Analysis of Deviance Table

Model	Log(likelihood)	# Param's	Deviance	Test d.f.	P-value
Full model	-2.70337	5			
Fitted model	-2.70337	2	2.79202e-007	3	1
Reduced model	-20.527	1	35.6472	4	<.0001

AIC: 9.40673

Goodness of Fit

Dose	Est._Prob.	Expected	Observed	Size	Scaled Residual
0.0000	0.0000	0.000	0.000	6.000	0.000
0.0100	0.0000	0.000	0.000	6.000	-0.000
0.1000	0.8334	5.000	5.000	6.000	-0.000
0.5000	1.0000	6.000	6.000	6.000	0.000
1.0000	1.0000	6.000	6.000	6.000	0.000

Chi^2 = 0.00 d.f. = 3 P-value = 1.0000

Benchmark Dose Computation

Specified effect = 0.1
 Risk Type = Extra risk
 Confidence level = 0.95
 BMD = 0.0526537
 BMDL = 0.00408511