

## Appendix 1

### **Power Simulations for Experimental Designs and Statistical Tests for Endocrine Disruption Experiments**

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The purpose of this simulation study was to explore the effects of number of replicates per treatment, allocation of replicates to treatments, number of subjects per replicate, number of treatment groups, and statistical test selection on the ability to detect statistically effects that might appear in an endocrine disruption study. The simulations were based on data collected in Phase 1 studies done in Japan and Germany for amphibian hindlimb length, but the results are general and apply to many measured responses. Additional simulations are under way and will be reported at a later date that specifically address developmental staging. Nevertheless, results from the simulation study reported here provide guidance for staging as well as hindlimb length and other measured response.

Two experimental Designs are considered. The first of these is allocation of equal numbers of replicates to each dose or treatment group. Thus the control and treatment groups all receive the same number of replicates and same number of subjects per replicate.

The second design assigns the same number of replicates to all treatment groups but more replicates to the control. Since the control group is used in every comparison of treatment to control, allocating more subjects to the control group than to the treatment groups will optimize power for a given total number of replicates. The total number of replicates is the same in both cases. Such a design also provides better estimation of the "natural" condition of experimental subjects and provides some protection against loss of a control replicate.

The statistical tests considered were the trend tests Williams and Jonckheere (both asymptotic and exact permutation) applied in a step-down fashion, and the pairwise tests Dunnett, Dunn, Mann-Whitney (asymptotic and exact permutation).

Some details of these tests and of the second design are given below. For simplicity, the term "dose" is used throughout as the exposure variable and does not imply any particular exposure mechanism.

At this stage of the simulation study, clear choices for experimental design and statistical test are evident. Further simulations, under way, are needed to translate these results into the power to detect changes of a specified size in continuous responses, such as hindlimb length, and ordinal responses, such as developmental stage. Also, tables rather than power curves are presented. When more complete simulations are available in a few weeks, power curves can and will be generated.

## **Summary of Findings**

### *Design Issues*

- 1) The minimum nominal number of replicates per treatment is recommended to be four. This is appropriate especially if a pairwise approach might be used, since for fewer than four replicates, the Dunn test is the only non-parametric test with positive power and its power properties are poor for fewer than four replicates.
- 2) Increased power is achieved through allocating replicates to control and treatment groups following the so-called square-root allocation rule.
- 3) It is clear from the simulations that for a given number of animals, much better power is obtained by putting fewer animals in more replicates than by putting more animals in fewer replicates.
- 4) The simulations also make clear that power is very clearly affected by test selection and allocation rule, power is largely determined by the number of replicates.

### *Test Selection Issues*

- 5) Where a monotone dose-response is expected biologically, the Williams or Jonckheere tests have approximately equal power, have superior power to pairwise tests, and are recommended, with a slight preference for Williams when the data are normally distributed and of homogeneous variance. For ordinal data, such as developmental stage, only the Jonckheere test is appropriate.
- 6) For fewer than four replicates per group, the Mann-Whitney test should not be used. With equal allocation of replicates, Mann-Whitney should not be used with fewer than five replicates.
- 7) For four replicates (or more) and square-root allocation of replicates, the Mann-Whitney test has good statistical properties when the normality and variance homogeneity requirements of Dunnett's test are not satisfied. Indeed, the Mann-Whitney powers exceed Dunnett test powers as the number of replicates per dose increases. As the number of replicates increases, the power of the Mann-Whitney test rivals that of Williams and Jonckheere.
- 8) The Dunnett test has lower power than either trend test under most conditions, but higher power than Dunn's test.
- 9) If a non-parametric test must be used, such as when no normalizing, variance stabilizing transform can be found or the data are inherently ordinal rather than quantitative, then the Dunn test is the choice for 3 or fewer replicates with square-root allocation and four or fewer replicates with equal allocation, while Mann-Whitney is the choice for 4 or more replicates per dose.

The improvement in the power properties of the tests from using the square-root allocation rule is real but generally less pronounced than the choice of test statistic or number of replicates. The square-root allocation affects some tests, such as Jonckheere, more than others. The square-root allocation should not be used when the nominal number of replicates per dose is 2 since this allocates only one replicate to each treatment group.

For experimental design, the square-root allocation rule with a nominal value of four replicates per dose is suggested. This puts three replicates in each positive dose group

and in the controls puts 7 replicates (for 4-dose experiment), 8 replicates (for 5 dose experiment), and 9 replicates (for 6 dose experiment). For equal allocation, five replicates per dose is recommended.

### **Details of Findings**

Tables 1, 4, and 7 show the comparisons of the seven tests under each choice of number of doses, number of replicates, and allocation rule. Tables 2, 5, and 8 compare the two allocation rules for each choice of number of doses, number of replicates, and test statistic. Tables 3, 6, and 9 show the effect of the number of replicates on each test statistic for each choice of number of doses and allocation rule.

### **Conditions of the Simulation**

Figure 1 shows the hindlimb length data from phase 1 reported by Japan and Germany plotted on a log scale (except for treatment of the zero concentration control) and shows for each country that the dose-response is roughly linear with respect to log(dose). The simulations accordingly followed a linear dose-response and Figure 2 shows one such simulation. Other dose-response shapes are of course possible and these shapes do affect the power properties of the test statistics. Several other dose-response shapes are being simulated and will be reported at a later date.

To understand the tables, the following will be helpful. The maximum effect simulated is stated and does not vary across the simulations regardless of number of doses. Then a linear dose-response trend is simulated running from zero effect to the specified maximum effect, with the effect at dose D proportional to the logarithm of dose, with the maximum dose always the same. Thus, at the high dose (be that 4, 5, or 6), the simulated mean response is the maximum effect. The effect at smaller doses will depend on the number of doses simulated. These effects are printed in the header for each table.

Once the number of doses and number of replicates per dose is selected and the mean or median response at each dose is determined as indicated above, a random normal number generator is used to populate each replicate and spread the replicates around the simulated mean response that is based on the within- and between-replicate variance observed in the two studies from phase 1.

The phase 1 studies used 5 doses, four positive treatment groups and one control. The simulation considered experiments with 4, 5, or 6 doses (including control). The phase 1 studies used two replicates of 25 subjects each. The simulation explored 2-8 replicates of 10, 15, 20, or 25 subjects per replicate. The hindlimb length measurements were found to be approximately normally distributed and the simulations generated random normal responses within each replicate. An analysis of variance (ANOVA) found within- and between-replicate variance and the simulations were given these same variance components as found in the Japan study. Finally, the same maximum effect as observed in the Japan study was simulated.

It should be observed that one effect of decreasing the number of subjects per replicate is to increase the variability of the replicate means simulated. Since the data are analyzed in

terms of replicate means, not individual subjects, decreasing the number of subjects per replicate has the effect of exploring increased variance. For example, cutting the number of replicates in half increases the standard deviation by roughly 40%. Subjects within the same replicate container in these endocrine disruption experiments are expected to be correlated rather than independent. It is for this reason that the statistical analysis treats the replicate mean (or median in the case of developmental stages) as the experimental unit rather than the individual.

An objective of the simulations was for the data from each simulated experiment to be analyzed by all seven tests. This would eliminate the remote chance that some simulation bias affects the comparisons. Since the two exact permutation tests require roughly 20 times the CPU time as the other five tests combined, in the interests of time, 1000-2000 experiments were simulated in which all seven tests were applied, while 5000-10000 experiments were simulated for the five other tests. In time, the number of experiments simulated with all seven tests applied will equal or exceed 5000. The standard error of the power estimates based on 5000 simulated experiments is expected to be no more than  $.5/\sqrt{5000}=.007$ , or less than 1%.

It is well understood that the number of replicates will affect the power of a statistical test, with higher power (i.e., greater ability to distinguish statistically the changes in mean or median response) associated with increased numbers of replicates. This simulation study will quantify the relationship between the number of replicates and the power of each statistical test examined.

#### **Statistical Tests Considered**

Seven statistical tests are explored. These are the Dunnett, Dunn, Williams, Jonckheere (or Jonckheere-Terpstra), and Mann-Whitney (or Wilcoxon). For the Jonckheere and Mann-Whitney tests, both the standard "asymptotic" and exact permutation versions of the tests are explored. The reason for this is that when only small numbers of replicates are present, the asymptotic or large-sample test statistics may not be accurate. Experience and theory both suggest exact permutation methods maybe required, especially when there are fewer than four replicates per concentration.

#### **Experimental Designs Considered**

The optimum allocation depends on the statistical test to be used. A widely used allocation rule was given by Dunnett (1955), which states that for a total of  $N$  replicates and  $k$  treatments to be compared to a common control, if the same number,  $n$ , of replicates are allocated to every treatment group, then the number,  $n_0$ , to allocate to the control to optimize power is determined by the so-called square-root rule. By this rule, the value of  $n$  is (the integer part of) the solution of the equation  $N = kn + n\sqrt{k}$ , and  $n_0 = N - kn$ . [It is almost equivalent to say  $n_0 = n\sqrt{k}$ .] This has been shown to optimize power for Dunnett's test. It is used, often without formal justification, for other pairwise tests, such as the Mann-Whitney and Fisher exact test. Williams (1972) showed that the square-root rule may be somewhat sub-optimal for his test and optimum power is achieved when  $\sqrt{k}$  in the above equation is replaced by something between  $1.1\sqrt{k}$  and  $1.4\sqrt{k}$ . For simplicity,

this minor modification is not followed in these simulations. The optimality of the square-root rule to other tests, such as Jonckheere-Terpstra and Cochran-Armitage has not been published in definitive form, but simulations (manuscript in preparation by J. W. Green) show that for the step-down Jonckheere-Terpstra test, power gains of up to 25% are possible under this rule compared to results from equal sample sizes. In all cases examined, the power is greater following this rule compared to equal sample sizes, where the total sample size is held constant. In the absence of definitive information on the Jonckheere-Terpstra and other tests, it is probably prudent to follow the square-root rule for pairwise and Jonckheere-Terpstra tests and either that or Williams' modification of the rule for other step-down procedures. The present simulation study does not show the square-root rule to be optimal, but it does demonstrate that this allocation rule improves the power properties of all tests considered (except under a few special circumstances, such as when the square-root rule puts only one replicate in each treatment group) and in every reasonable design.

#### **Discussion of Tests**

The Dunn, Mann-Whitney, and Jonckheere tests are non-parametric tests based on ranking the data by order of magnitude. Such tests are required when the data do not meet the requirements of the usual parametric procedures, which for the Dunnett and Williams tests are within-dose normality and variance homogeneity, plus independence of replicate means (or medians). Often, a simple transformation of the response will resolve a problem with normality or variance homogeneity. In other cases, no normalizing, variance stabilizing transformation can be found. Developmental stage is an inherently non-normal response and one of these non-parametric tests would be appropriate. It would not be appropriate, for example, to analyze developmental stage by use of a chi-square statistic based on a multinomial distribution, since such a test can be significant when the only difference between two dose groups is an increase or decrease in variation. The intent is to detect an increase in median developmental stage, not a change in spread of the stages.

#### **A Comment on Parametric and Non-Parametric Tests**

One often hears criticism of non-parametric tests on the grounds that they have lower power and one also hears claims that parametric tests are always to be preferred. This simulation study is an unbiased assessment of these claims in this specific context and for the specific tests considered. In fact, the non-parametric Jonckheere test has very good power properties in general and the Mann-Whitney does also provided there are four or more replicates per dose.

#### **Pairwise versus Trend Tests**

In addition to the division of tests between parametric and non-parametric, the Williams and Jonckheere tests are step-down trend tests, while the others are pairwise tests.

The hypothesis that is tested in determining the NOEC for a toxicological experiment reflects the risk assessment question and the assumptions that are made concerning the underlying characteristics, or statistical model, of the responses being analyzed (e.g., does the response increase in an orderly (i.e., monotone) way with increasing toxicant dose or

concentration?). The statistical test that is used depends on the hypothesis tested (e.g., are responses in all groups equal?), the associated statistical model, and the distribution of the values (e.g., are data normally distributed?). Thus, it is necessary to understand the question to be answered and to translate this question into appropriate null and alternative hypotheses before selecting the test procedure.

The simplest statistical model generally used in hypothesis testing assumes only that the distributions of responses within these populations are identical except for a location parameter (e.g., the mean or median of the distribution of values from each group). Another statistical model that is often used assumes that there is a trend in the response that is associated with increasing exposure. Each of these models suggests a set of hypotheses that can be tested to determine whether the model is consistent with the data. These two types of hypotheses can further be expressed as 1-sided or 2-sided. The discussion below is developed in terms of population means, but applies equally to hypotheses concerning population medians. The most basic hypothesis (in 1-sided form) can be stated as follows:

$H_0 : \mu_0 = \mu_1 = \mu_2 = \dots = \mu_k$  vs.  $H_1 : \mu_0 > \mu_i$  for at least one  $i$ , (Model 1)

where  $\mu_i$ ,  $i=0, 1, 2, 3, \dots, k$  denote the means of the control and test *populations*, respectively.

Thus, one tests the null hypothesis of no differences among the population means against the alternative that at least one population mean is smaller than the control mean. There is no investigation of differences among the treatment means, only whether treatment means differ from the control mean. The one-sided hypothesis is appropriate when an effect in only one direction is a concern. The direction of the inequality in the above alternative hypothesis (i.e. in  $H_1 : \mu_0 > \mu_i$ ) would be appropriate if a decrease in the endpoint was a concern but an increase was not (for instance, if an exposure was expected to induce infertility and reduce number of offspring). If an increase in the endpoint was the only concern, then the direction of the inequality would be reversed.

The conceptual difference between these two types of tests is as follows. Pairwise tests compare each treatment group to the control independent of all the other treatment groups. It is entirely possible, for example, for such a test to find a significant difference between dose 2 and the control but not find a significant difference between dose 3 or 4 and the control. A trend test cannot produce such a result. In comparing dose 4 to the control, all the doses between dose 4 and the control are taken into account. If no significant trend is found with all doses present, then no further testing is done and the high dose is the NOAEL. On the other hand, if the trend with all doses included is significant, then the high dose is an effects level. One then omits the high dose group and repeats the test with the remaining dose groups. If the trend test with the dose groups is significant, the test is repeated with the current high dose group omitted. This process continues until the first time the trend test is not significant. At that stage, testing stops and the NOAEL is the highest dose remaining. Such a procedure is justified on the grounds that biologically, one expects a toxic effect to manifest itself more strongly as the dose increases. That is, one expects a monotonic dose-response relationship. Should

this not be expected biologically, then a trend test is not appropriate and pairwise methods should be used.

If no assumption is made about the relationships among the treatment groups and control (e.g., no trend is assumed), the test statistics will be based on comparing each treatment to the control, independent of the other treatments. Many tests have been developed for this approach, mostly for experiments in which treatments are qualitatively different, as, for example, in comparing various new therapies or drug formulations to a standard.

In toxicology, the treatment groups generally differ only in the exposure concentration (or dose) of a single chemical. It is further often true that biology suggests that if the chemical is toxic, then as the level of exposure is increased, the magnitude effect will tend to increase. Depending on what response is measured, the effect of increasing exposure may show up as an increase or as a decrease in the measured response, but not both. The statistical model underlying this biological expectation is what will be called a trend model or a model assuming monotonicity of the population means:

$$\mu_0 \geq \mu_1 \geq \mu_2 \geq \mu_3 \geq \dots \geq \mu_k \text{ (or with inequalities reversed)}$$

The null and alternative hypotheses can then be stated as

$$H_{02} : \mu_0 = \mu_1 = \mu_2 = \dots = \mu_k \text{ vs } H_{12} : \mu_0 \geq \mu_1 \geq \mu_2 \geq \mu_3 \geq \dots \geq \mu_k, \text{ with } \mu_0 > \mu_k.$$

Note that  $\mu_0 > \mu_k$  is equivalent, under the alternative, to  $\mu_0 > \mu_i$  for at least one  $i$ . If this monotone model is accepted as representing the true responses of test organisms to exposure to toxicants, it is not possible for, say,  $\mu_3$  to be smaller than  $\mu_0$  and  $\mu_6$  not to be smaller.

Under the trend model and tests designed for that model, if tests of hypotheses  $H_{02}$  vs.  $H_{12}$  reveal that  $\mu_3$  is different from  $\mu_0$ , but  $\mu_2$  is not, the NOEC has been determined (i.e. it is the test concentration associated with  $\mu_2$ ), and there is no need to test whether  $\mu_1$  differs from  $\mu_0$ . Also, finding that  $\mu_3$  differs from  $\mu_0$  implies that a significant trend exists across the span of doses including  $\mu_0$  and  $\mu_3$ , the span including  $\mu_0$  and  $\mu_4$ , and so on. For the majority of toxicological studies, a test of the trend hypothesis based on model (2) is consistent with the basic expectations for a model for dose-response. In addition, statistical tests for trend tend to be more powerful than alternative non-trend tests, and should be the preferred tests if they are applicable. Thus, a necessary early step in the analysis of results from a study is to consider each endpoint, decide whether a trend model is appropriate, and then choose the initial statistical test based on that decision. Only after it is concluded trend is not appropriate do specific pairwise comparisons make sense to illuminate sources of variability.

#### Adjustments to P-values for Multiple Tests

A common problem with doing several related statistical tests is that each test carries with it a non-zero chance of declaring a difference between treatment and control to be significant when in fact there is no real treatment effect. Typically, one accepts a 5% chance of error of this type. If numerous tests are done, each with a 5% chance of incorrectly declaring an effect to be significant when no true difference exists, then the chance that at least one of these tests falsely declaring a significant effect is higher than 5%, sometimes much higher. As a consequence, some adjustment is usually made to

control the overall chance of at least one of the many tests being wrong. The Dunnett test has a built-in adjustment. The Williams and Jonckheere tests are applied in step-down fashion, with a significant result at one stage a requirement to proceed to the next stage. Consequently, each step in applying Williams or Jonckheere is done at the same 5% level and in this particular application, the overall false positive rate remains fixed at 5%. The Dunn and Mann-Whitney tests have no built-in multiplicity adjustment. There are several adjustment schemes. The Bonferroni-Holms adjustment is a considerable improvement over the older Bonferroni adjustment and is the method used in these simulations.

*Pairwise comparison procedures* amount to performing all possible comparisons of treatment groups to the control. Multiple comparisons to the control may be made, but there is no ordered set of hypotheses to test, and no use of the sequence of outcomes in deciding which comparisons to make. Examples of the single-step approach include the use of the Fisher's exact test, the Mann-Whitney, Dunnett and Dunn tests. Since many comparisons to the control are made, some adjustment must be made for the number of such comparisons to keep the family-wise error (FWE) rate at a fixed level, generally 0.05. With tests that are inherently single comparison tests, such as Fisher's exact and Mann-Whitney, a Bonferroni adjustment can be made: a study with  $k$  treatment levels would be analyzed by performing the pair-wise comparisons of each of the treatment groups to the control group, each performed at a significance level of  $\alpha/k$  instead of  $\alpha$ . (This is the Bonferroni adjustment.) Equivalently, the calculated p-value ignoring multiplicities is multiplied by  $k$ . That is,  $p_i^b = k * p_i$ . The Bonferroni adjustment is generally overly conservative, especially for large  $k$ . Modifications reduce the conservatism while preserving the FWE at 0.05 or less.

For the Holm modification of the Bonferroni adjustment, arrange the  $k$  unadjusted p-values for all comparisons of treatments to control in rank order, i.e.,  $p_{(1)} \leq p_{(2)} \leq p_{(3)} \leq \dots \leq p_{(k)}$ . Beginning with  $p_{(1)}$ , compare  $p_{(1)}$  with  $\alpha/(k-i+1)$ , stopping at the first non-significant comparison. If the smallest  $i$  for which  $p_{(i)}$  exceeds  $\alpha/(k-i+1)$  is  $i=j$ , then all comparisons with  $i>j$  are judged non-significant without further comparisons. It is helpful (Wright (1992)) to report adjusted p-values rather than the above comparisons. Thus, report  $p_{(1)}^* = p_{(1)} * (k-i+1)$  and then compare each adjusted p-value to  $\alpha$ . The table below illustrates the advantage of the Bonferroni-Holm method. In this hypothetical example, only the comparison of treatment 4 with the control would be significant if the Bonferroni adjustment is used, whereas all comparisons except the comparison of the Control with treatment 1 would be significant if the Bonferroni-Holm adjustment is used.

Comparison	Unadjusted p value	Bonferroni-Holm Adjusted p value $p_{(i)}^*$	Bonferroni Adjusted p values $P_i^b$
Control – Treatment 4	$p_{(1)} = 0.002$	$0.002 * 4 = 0.008$	$0.002 * 4 = 0.008$
Control – Treatment 2	$p_{(2)} = 0.013$	$0.013 * 3 = 0.039$	$0.013 * 4 = 0.052$
Control – Treatment 3	$p_{(3)} = 0.020$	$0.020 * 2 = 0.040$	$0.02 * 4 = 0.08$
Control – Treatment 1	$p_{(4)} = 0.310$	$0.310 * 1 = 0.310$	$0.310 * 4 = 1$

*Step-down procedures* are generally preferred where they are applicable. All step-down procedures discussed are based on a sequential process consisting of testing an ordered set of hypotheses concerning means, ranks, or trend. A step-down procedure based on trend (for example) works as follows: First, the hypothesis that there is no trend in response with increasing dose is tested when the control and all dose groups are included in the test. Then, if the test for trend is significant, the high dose group is dropped from the data set, and the hypothesis that there is no trend in the reduced data set is tested. This process of dropping treatment groups and testing is continued until the first time the trend test is non-significant. The highest dose in the reduced data set at that stage is then declared to be the NOEC. Distinguishing features of step-down procedures are that the tests of hypothesis must be performed in a given order, and that the outcome of each hypothesis test is evaluated before deciding whether to test the next hypothesis in the ordered sequence of hypotheses. It is these two aspects of these procedures that account for controlling the family-wise error (FWE) rate.

A step-down method typically uses a critical level larger than that used in single-step procedures, and seeks to limit the number of comparisons that need to be made. Indeed, the special class of "fixed-sequence" tests described below fix the critical level at 0.05 for each comparison but bound the FWE rate at 0.05. Thus, step-down methods are generally preferable to the single-step methods as long as the response means are monotonic. Tests based on trend are logically consistent with the anticipated monotone pattern of responses in toxicity tests. Step-down procedures make use of this ordered alternative by ordering the tests of hypotheses. This minimizes the number of comparisons that need to be made, and in all the methods discussed here, a trend model is explicitly assumed (and tested) as a part of the procedure.

Procedures that employ step-down trend tests have more power than procedures that rely on multiple pairwise comparisons when there is a monotone dose-response because they make more use of the biology and experimental design being analyzed. When there is a monotone dose-response, procedures that compare single treatment means or medians against the control, independent of the results in other treatments (i.e. single-step procedures), ignore important and relevant information, and suffer power loss as a result. The trend models used in the step-down procedures do not assume a particular precise mathematical relationship between dose and response, but rather use only monotonicity of the dose-response relationship. The underlying statistical model assumes a monotone dose-response in the *population* means, not the *observed* means.

Rejection of the null hypothesis (i.e., rejecting the hypothesis that all group means, or medians, or distributions are equal) in favor of the stated alternative implies that the high dose is significantly different from the control. The same logic applies at each stage in the step-down application of the test to imply, whenever the test is significant, that the high dose remaining at that stage is significantly different from the control. These tests are all applied in a 1-sided manner with the direction of the alternative hypothesis always the same. Moreover, this methodology is general, and applies to any legitimate test of the stated hypotheses under the stated model. That is, one can use this fixed-sequence

approach with the Cochran-Armitage test on quantal data, the Jonckheere-Terpstra or Williams or Brown-Forsythe tests of trend on continuous data. Other tests of trend can also be used in this manner.

#### The Power Concept

In addition to controlling the risk of a false positive, one must also be concerned with the risk of a false negative, that is, the chance of not declaring a difference between treatment and control to be significant when in fact there is a real difference of biological importance. It is convenient to refer to the chance of correctly declaring significant a treatment effect instead of the chance of not finding such an effect significant. These two chances are complementary, that is, they always add to 100%. The chance of correctly declaring significant a real treatment effect is the power of the test to detect that difference and varies with the size of the true effect.

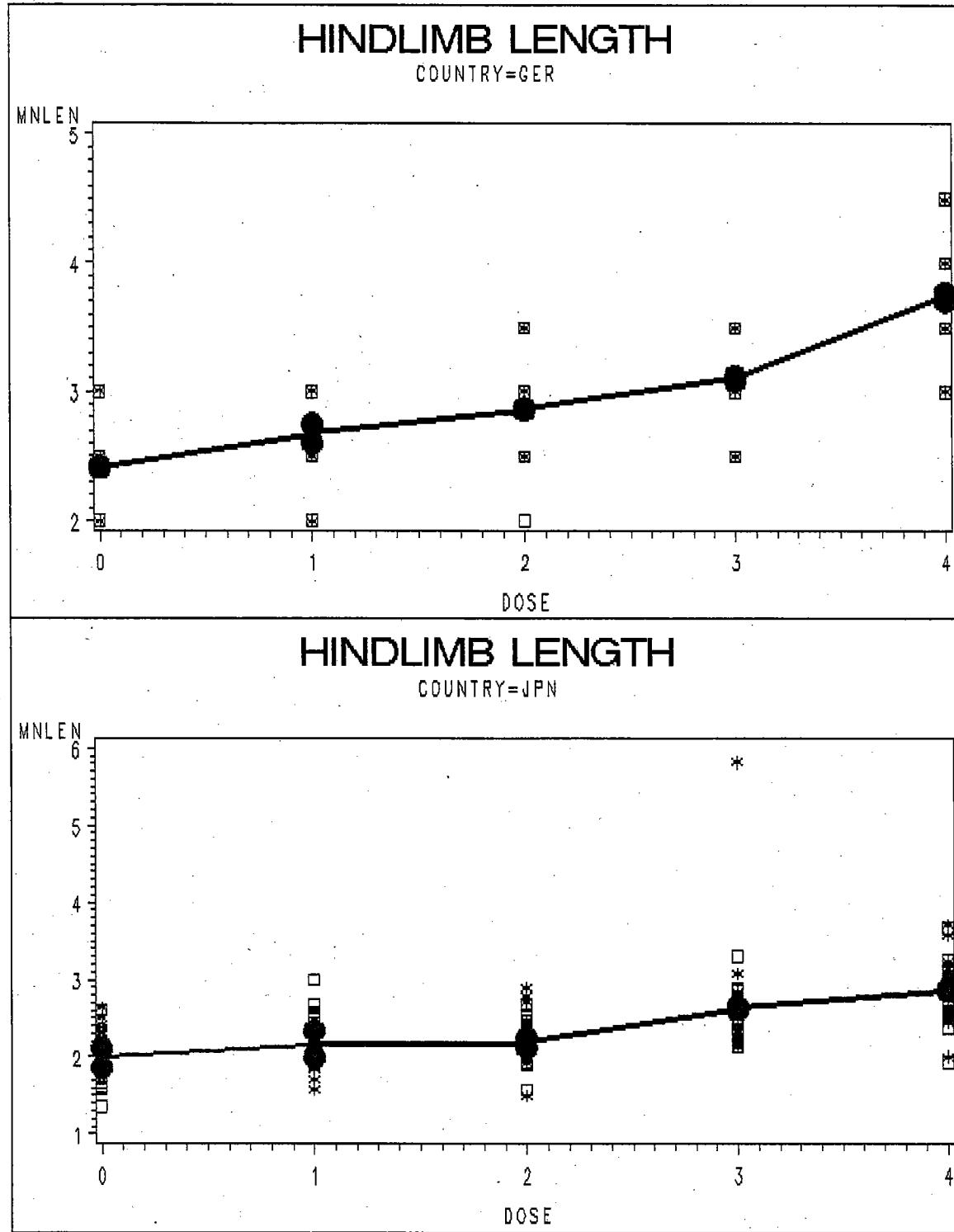


Figure 1

The red dots are replicate means and the line joins the treatment means.

# SIMULATED HINDLIMB LENGTH

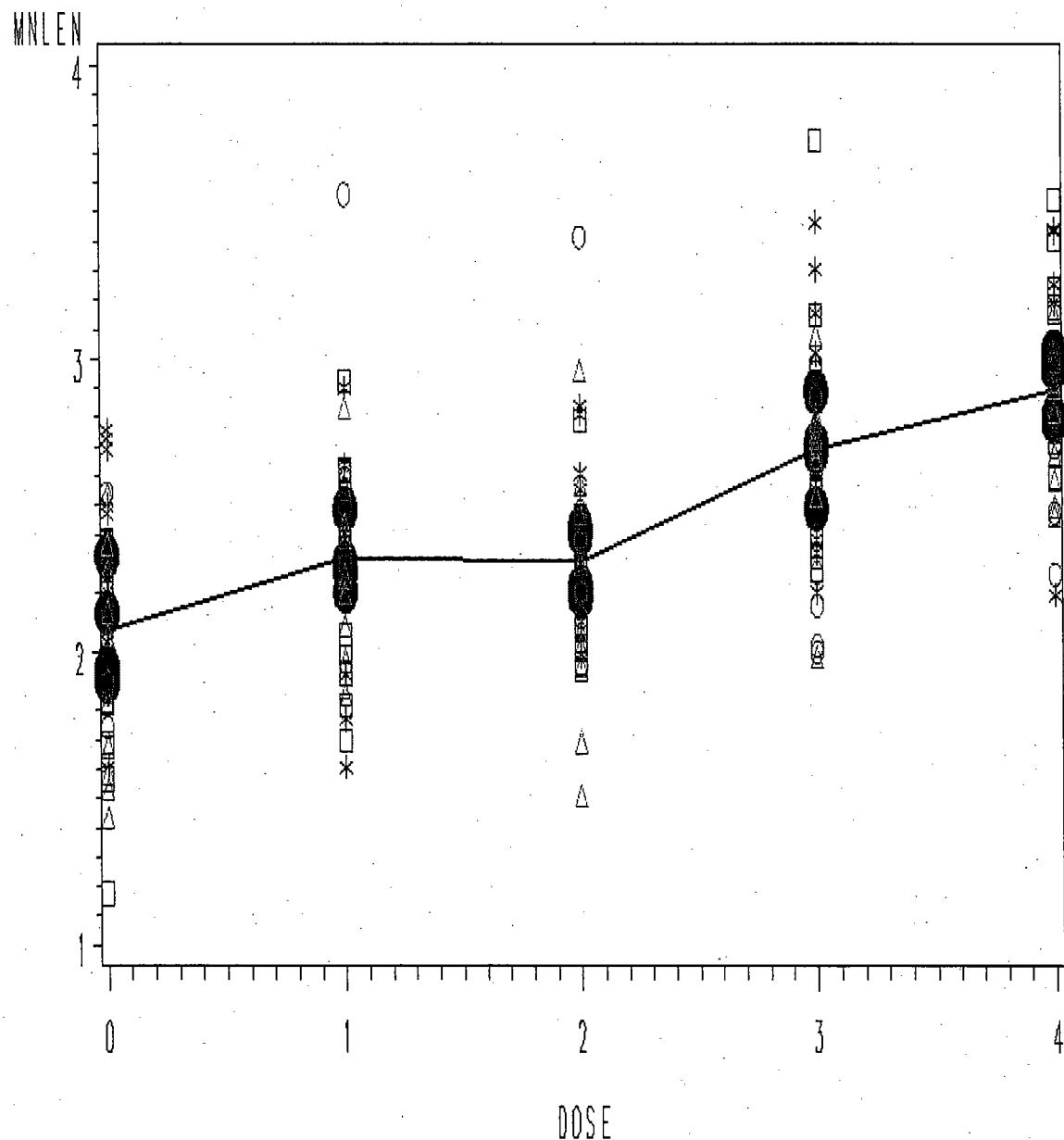


Figure 2

This simulates four replicates of 10 subjects each with standard allocate rule. Red dots are replicate means. The line joins the treatment means.

**Table 1 Comparison of Tests for 4 Dose Experiments**  
**POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE**  
**TREND RELATIVE TO LOG(DOSE)**  
**MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754**

REPS	SAMP SIZE	SQRTRULE TEST	NOAEL			
			0	1	2	3
2	NO	DUNN		74	26	
		DUNNETT	21	52	25	4
		JONCKH		75	25	1
		JONCKHX		75	25	1
		MW			100	
		MWX			100	
	YES	WILLIAMS	34	52	14	1
		DUNN			100	
		DUNNETT	12	46	34	9
		JONCKH		86	12	3
		JONCKHX		62	31	7
		MW			100	
15	NO	MWX			100	
		WILLIAMS	30	51	18	2
		DUNN			83	17
		DUNNETT	26	54	19	1
		JONCKH		82	17	0
		MW			100	
	YES	WILLIAMS	42	49	9	0
		DUNN			100	
		DUNNETT	19	49	27	4
		JONCKH		90	8	2
		MW			100	
		WILLIAMS	38	47	14	0
20	YES	DUNN			100	
		DUNNETT	25	51	22	2
		JONCKH		91	7	2
		MW			100	
		WILLIAMS	42	48	9	0
		DUNN			87	14
	NO	DUNNETT	33	54	12	1
		JONCKH		87	14	0
		JONCKHX		88	13	0
		MW			100	
		MWX			100	
		WILLIAMS	50	45	6	0
25	YES	DUNN			100	
		DUNNETT	25	51	21	3
		JONCKH		92	7	1
		JONCKHX		73	24	4
		MW			100	
		MWX			100	
	NO	WILLIAMS	43	48	8	0
		DUNN		23	77	1
		DUNNETT	43	53	5	0
		JONCKH		56	43	2
		JONCKHX		28	68	4
		MW			100	
3	10	MWX			3	97
		WILLIAMS	59	40	2	
		DUNN		17	82	2
		DUNNETT	41	55	5	
		JONCKH		66	31	4
		JONCKHX		49	48	3
	NO	MW			100	
		MWX			14	86
		WILLIAMS	59	40	1	

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**TREND RELATIVE TO LOG(DOSE)**  
**MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754**

REPS	SAMPLESIZE	SQRTRULE	TEST	NOAEL			
				0	1	2	3
15	NO	WILLIAMS	DUNN	20	80	0	
			DUNNETT	53	46	2	
			JONCKH	61	38	1	
			MW			100	
			WILLIAMS	68	32	1	
	YES	DUNN	DUNN		16	83	1
			DUNNETT	49	49	2	
			JONCKH	73	25	2	
			MW			100	
			WILLIAMS	68	32	1	
20	NO	WILLIAMS	DUNN		20	80	
			DUNNETT	55	44	1	
			JONCKH	63	36	1	
			MW			100	
			WILLIAMS	71	29	0	
	YES	DUNN	DUNN		14	85	1
			DUNNETT	52	46	2	
			JONCKH	76	22	2	
			MW			100	
			WILLIAMS	70	30	1	
25	NO	WILLIAMS	DUNN		21	79	0
			DUNNETT	60	40	0	
			JONCKH	68	32	0	
			JONCKHX	34	66	1	
			MW			100	
	YES	MWX	MWX		3	97	
			WILLIAMS	75	25	0	
			DUNN		13	87	0
			DUNNETT	57	42	1	
			JONCKH	78	20	2	
4	10	NO	JONCKHX	62	37	2	
			MW			100	
			MWX		20	81	
			WILLIAMS	74	26	0	
			DUNN		70	31	
	YES	WILLIAMS	DUNNETT	59	42	1	
			JONCKH	72	28	0	
			JONCKHX	57	43	0	
			MW	55	38	7	0
			MWX	56	38	7	0
15	NO	WILLIAMS	WILLIAMS	72	29	0	
			DUNN	0	84	16	
			DUNNETT	60	40	1	
			JONCKH	75	25	1	
			JONCKHX	65	35	1	
	YES	MW	MW	64	34	2	0
			MWX	64	34	2	0
			WILLIAMS	76	25	0	
			DUNN		77	23	
			DUNNETT	67	33	0	

**Table 1 Comparison of Tests for 4 Dose Experiments**  
**POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE**  
**TREND RELATIVE TO LOG(DOSE)**  
**MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754**

REPS	SAMP SIZE	SQRTRULE	TEST	NOAEL		
				0	1	2
4	20	NO	DUNN	81	19	
			DUNNETT	73	27	0
			JONCKH	82	18	0
			MW	68	30	2
			WILLIAMS	84	16	
	YES		DUNN	91	9	
			DUNNETT	74	26	0
			JONCKH	86	15	
			MW	76	23	1
			WILLIAMS	86	14	
25	NO		DUNN	84	16	
			DUNNETT	77	23	0
			JONCKH	85	15	
			JONCKHX	73	28	
			MW	72	27	2
	YES		MWX	72	27	2
			WILLIAMS	87	13	
			DUNN	93	7	
			DUNNETT	76	24	
			JONCKH	86	14	
5	10	NO	JONCKHX	79	22	0
			MW	78	22	0
			MWX	79	22	0
			WILLIAMS	87	13	
			DUNN	92	8	
	YES		DUNNETT	70	30	0
			JONCKH	79	22	
			JONCKHX	76	24	
			MW	78	21	1
			MWX	76	23	1
15	NO		WILLIAMS	83	17	
			DUNN	2	96	3
			DUNNETT	74	26	0
			JONCKH	85	16	0
			JONCKHX	78	22	0
	YES		MW	78	22	1
			MWX	78	22	0
			WILLIAMS	86	14	
			DUNN	97	3	
			DUNNETT	79	21	
20	NO		JONCKH	85	15	
			MW	85	15	0
			WILLIAMS	88	12	
			DUNN	1	98	1
			DUNNETT	81	19	0
	YES		JONCKH	90	10	
			MW	84	15	0
			WILLIAMS	90	10	
			DUNN	98	2	
			DUNNETT	85	15	

**Table 1 Comparison of Tests for 4 Dose Experiments**  
**POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE**  
**TREND RELATIVE TO LOG(DOSE)**  
**MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754**

REPS	SAMP SIZE	SQRTRULE TEST	NOAEL			
			0	1	2	3
25	NO	DUNN	99	1		
		DUNNETT	87	13		
		JONCKH	91	9		
		JONCKHX	90	10		
		MW	91	9	0	
	YES	MWX	90	10	0	
		WILLIAMS	94	6		
		DUNN	0	99	0	
		DUNNETT	89	11		
		JONCKH	93	7		
6	10	JONCKHX	90	10		
		MW	90	10	0	
		MWX	91	10		
		WILLIAMS	95	5		
		DUNN	0	98	2	
	YES	DUNNETT	81	19		
		JONCKH	86	14		
		JONCKHX	86	15		
		MW	86	14	0	
		MWX	85	15	0	
15	NO	WILLIAMS	89	11		
		DUNN	4	96	0	
		DUNNETT	82	18		
		JONCKH	91	9		
		MW	88	12	0	
	YES	WILLIAMS	91	9		
		DUNN	0	100	0	
		DUNNETT	88	12		
		JONCKH	91	9		
		MW	91	9		
20	NO	WILLIAMS	94	6		
		DUNN	3	97	0	
		DUNNETT	90	10		
		JONCKH	95	5		
		MW	93	7		
	YES	WILLIAMS	95	5		
		DUNN	0	100	0	
		DUNNETT	91	9		
		JONCKH	93	7		
		MW	93	7		
25	NO	WILLIAMS	95	5		
		DUNN	2	98	0	
		DUNNETT	93	7		
		JONCKH	95	5		
		JONCKHX	95	6		
	YES	MW	95	5		
		MWX	95	6		
		WILLIAMS	97	3		
		DUNN	2	98	0	
		DUNNETT	94	6		

**Table 1 Comparison of Tests for 4 Dose Experiments**  
**POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE**  
**TREND RELATIVE TO LOG(DOSE)**  
**MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754**

REPS	SAMP SIZE	SQRTRULE TEST	NOAEL			
			0	1	2	3
7	10	NO	DUNN	1	99	0
			DUNNETT	87	13	
			JONCKH	91	9	
			MW	91	9	0
			WILLIAMS	93	7	
	YES	YES	DUNN	17	83	0
			DUNNETT	87	13	
			JONCKH	92	8	
			MW	92	8	
			WILLIAMS	95	6	
15	NO	NO	DUNN	0	100	0
			DUNNETT	93	7	
			JONCKH	96	4	
			MW	96	4	
			WILLIAMS	97	3	
	YES	YES	DUNN	17	83	
			DUNNETT	93	7	
			JONCKH	96	4	
			MW	96	4	
			WILLIAMS	97	3	
20	NO	NO	DUNN	0	100	0
			DUNNETT	95	5	
			JONCKH	97	3	
			MW	97	3	
			WILLIAMS	98	2	
	YES	YES	DUNN	16	84	
			DUNNETT	95	5	
			JONCKH	97	3	
			MW	97	3	
			WILLIAMS	98	2	
25	NO	NO	DUNN	0	100	0
			DUNNETT	96	4	
			JONCKH	98	2	
			JONCKHX	98	3	
			MW	98	2	
	YES	YES	MWX	98	3	
			WILLIAMS	99	1	
			DUNN	16	84	
			DUNNETT	97	3	
			JONCKH	98	2	
8	10	NO	JONCKHX	98	2	
			MW	98	2	
			MWX	98	2	
			WILLIAMS	99	1	
			DUNN	2	98	0
	YES	YES	DUNNETT	92	8	
			JONCKH	94	6	
			MW	93	8	
			WILLIAMS	96	4	
			DUNN	26	74	

**Table 1 Comparison of Tests for 4 Dose Experiments**  
POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE  
TREND RELATIVE TO LOG(DOSE)  
MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

REPS	SAMP SIZE	SQR RULE TEST	NOAEL			
			0	1	2	3
8	15	NO	DUNN	2	98	
			DUNNETT	96	4	
			JONCKH	98	2	
			MW	97	3	
			WILLIAMS	99	1	
	YES	YES	DUNN	30	70	
			DUNNETT	96	4	
			JONCKH	98	2	
			MW	97	3	
			WILLIAMS	98	2	
20	NO	NO	DUNN	1	99	
			DUNNETT	97	3	
			JONCKH	98	2	
			MW	97	3	
			WILLIAMS	99	1	
	YES	YES	DUNN	35	65	
			DUNNETT	98	2	
			JONCKH	99	1	
			MW	98	2	
			WILLIAMS	99	1	
25	NO	NO	DUNN	1	99	
			DUNNETT	98	2	
			JONCKH	99	1	
			JONCKHX	99	2	
			MW	99	1	
	YES	YES	MWX	99	2	
			WILLIAMS	99	1	
			DUNN	37	64	
			DUNNETT	98	2	
			JONCKH	99	1	
			JONCKHX	99	1	
			MW	99	1	
			MWX	99	1	
			WILLIAMS	100	1	

**Table 2**  
**Comparison of Allocation Rules for 4 Dose Experiments**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND  
 RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST DUNN	REPS	SAMP SIZE	SQRTRULE	NOAEL			
				0	1	2	3
2	10	NO			74	26	
		YES				100	
		15	NO		83	17	
			YES			100	
		20	YES			100	
	15	25	NO		87	14	
			YES			100	
		10	NO	23	77	1	
			YES	17	82	2	
		15	NO	20	80	0	
3	10		YES	16	83	1	
		20	NO	20	80		
			YES	14	85	1	
		25	NO	21	79	0	
			YES	13	87	0	
	15	10	NO	70	31		
			YES	0	84	16	
		15	NO	77	23		
			YES	0	87	13	
		20	NO	81	19		
4	10		YES	91	9		
		25	NO	84	16		
			YES	93	7		
		10	NO	92	8		
			YES	2	96	3	
	15	15	NO	97	3		
			YES	1	98	1	
		20	NO	98	2		
			YES	1	99	1	
		25	NO	99	1		
5	10		YES	0	99	0	
		10	NO	0	98	2	
			YES	4	96	0	
		15	NO	0	100	0	
			YES	3	97	0	
	15	20	NO	0	100	0	
			YES	2	98	0	
		25	NO	0	100	0	
			YES	2	98	0	
		20	NO	1	99	0	
6	10		YES	17	83	0	
		10	NO	0	100	0	
			YES	4	96	0	
		15	NO	0	100	0	
			YES	3	97	0	
	15	20	NO	0	100	0	
			YES	2	98	0	
		25	NO	0	100	0	
			YES	2	98	0	
		20	NO	1	99	0	
7	10		YES	17	83	0	
		10	NO	0	100	0	
			YES	17	83	0	
		15	NO	0	100	0	
			YES	17	83	0	
	15	20	NO	0	100	0	
			YES	16	84		
		25	NO	0	100	0	
			YES	16	84		
		20	NO	0	100	0	
8	10		YES	16	84		
		10	NO	2	98	0	
			YES	26	74		
		15	NO	2	98		
			YES	30	70		
	15	20	NO	1	99		
			YES	35	65		
		25	NO	1	99		
			YES	37	64		

**Table 2**  
**Comparison of Allocation Rules for 4 Dose Experiments**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST	REPS	SAMP SIZE	SQR RULE	NOAEL			
				0	1	2	3
DUNNETT	2	10	NO	21	52	25	4
			YES	12	46	34	9
		15	NO	26	54	19	1
			YES	19	49	27	4
			YES	25	51	22	2
	3	20	NO	33	54	12	1
			YES	25	51	21	3
		25	NO	43	53	5	0
			YES	41	55	5	
			NO	53	46	2	
4	4	10	YES	49	49	2	
			NO	55	44	1	
		15	YES	52	46	2	
			NO	60	40	0	
			YES	57	42	1	
	5	20	NO	59	42	1	
			YES	60	40	1	
		25	NO	67	33	0	
			YES	69	31	0	
			NO	73	27	0	
6	6	10	YES	74	26	0	
			NO	77	23	0	
		15	YES	76	24		
			NO	70	30	0	
			YES	74	26	0	
	7	20	NO	79	21		
			YES	81	19	0	
		25	NO	85	15		
			YES	85	15	0	
			NO	87	13		
8	8	10	YES	89	11		
			NO	81	19		
		15	YES	82	18		
			NO	88	12		
			YES	90	10		
	9	20	NO	91	9		
			YES	93	7		
		25	NO	93	7		
			YES	94	6		
			NO	87	13		
10	10	10	YES	87	13		
			NO	93	7		
		15	YES	93	7		
			NO	95	5		
			YES	95	5		
	15	20	NO	96	4		
			YES	97	3		
		25	NO	97	3		
			YES	98	2		
			NO	98	2		
			YES	98	2		

**Table 2**  
**Comparison of Allocation Rules for 4 Dose Experiments**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST JONCKH	REPS	SAMP SIZE	SQRTRULE	NOAEL			
				0	1	2	3
2	10	NO		75	25	1	
		YES		86	12	3	
		15	NO	82	17	0	
		YES		90	8	2	
		20	YES	91	7	2	
		25	NO	87	14	0	
	15	YES		92	7	1	
		10	NO	56	43	2	
		YES		66	31	4	
		20	NO	61	38	1	
		YES		73	25	2	
		25	NO	63	36	1	
3	10	YES		76	22	2	
		25	NO	68	32	0	
		YES		78	20	2	
		15	NO	72	28	0	
		YES		75	25	1	
		20	NO	78	22	0	
	15	YES		81	19	0	
		25	NO	82	18	0	
		YES		86	15		
		20	NO	85	15		
		YES		86	14		
		25	NO	79	22		
4	10	YES		85	16	0	
		15	NO	85	15		
		YES		90	10		
		20	NO	89	11		
		YES		92	8		
		25	NO	91	9		
	15	YES		93	7		
		10	NO	86	14		
		YES		91	9		
		20	NO	91	9		
		YES		95	5		
		25	NO	93	7		
5	10	YES		96	4		
		25	NO	95	5		
		YES		97	3		
		15	NO	91	9		
		YES		92	8		
		20	NO	96	4		
	15	YES		96	4		
		25	NO	97	3		
		YES		97	3		
		20	NO	98	2		
		YES		98	2		
		25	NO	98	2		
6	10	YES		99	1		
		25	NO	95	5		
		YES		97	3		
		15	NO	96	4		
		YES		95	5		
		20	NO	93	7		
	15	YES		96	4		
		25	NO	95	5		
		YES		97	3		
		20	NO	98	2		
		YES		98	2		
		25	NO	98	2		
7	10	YES		99	1		
		25	NO	99	1		
		YES		99	1		
		15	NO	96	4		
		YES		96	4		
		20	NO	97	3		
	15	YES		97	3		
		25	NO	98	2		
		YES		98	2		
		20	NO	98	2		
		YES		99	1		
		25	NO	99	1		
8	10	YES		99	1		
		25	NO	99	1		
		YES		99	1		
		15	NO	98	2		
		YES		98	2		
		20	NO	98	2		
	15	YES		99	1		
		25	NO	99	1		
		YES		99	1		
		20	NO	99	1		
		YES		99	1		
		25	NO	99	1		

**Table 2**  
**Comparison of Allocation Rules for 4 Dose Experiments**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST	REPS	SAMP SIZE	SQRTRULE	NOAEL			
				0	1	2	3
JONCKHX	2	10	NO	75	25	1	
			YES	62	31	7	
		25	NO	88	13	0	
			YES	73	24	4	
	3	10	NO	28	68	4	
			YES	49	48	3	
		25	NO	34	66	1	
			YES	62	37	2	
4	4	10	NO	57	43	0	
			YES	65	35	1	
		25	NO	73	28		
			YES	79	22	0	
	5	10	NO	76	24		
			YES	78	22	0	
		25	NO	90	10		
			YES	90	10		
6	6	10	NO	86	15		
			NO	95	6		
		25	YES	96	5		
			NO	98	3		
	7	25	YES	98	2		
			NO	99	2		
		25	YES	99	1		
			NO	100			
MW	2	10	NO	100			
			YES	100			
		15	NO	100			
			YES	100			
		20	NO	100			
			YES	100			
		25	NO	100			
			YES	100			
	3	10	NO	100			
			YES	100			
		15	NO	100			
			YES	100			
		20	NO	100			
			YES	100			
		25	NO	100			
			YES	100			
4	4	10	NO	55	38	7	0
			YES	64	34	2	0
		15	NO	64	32	4	0
			YES	71	27	1	0
		20	NO	68	30	2	
			YES	76	23	1	
		25	NO	72	27	2	
			YES	78	22	0	0
	5	10	NO	78	21	1	
			YES	78	22	1	
		15	NO	85	15	0	
			YES	84	15	0	
		20	NO	89	11	0	
			YES	87	13	0	
		25	NO	91	9	0	
			YES	90	10	0	

**Table 2**  
**Comparison of Allocation Rules for 4 Dose Experiments**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST	REPS	SAMP SIZE	SQRTRULE	NOAEL			
				0	1	2	3
6	10	NO		86	14	0	
		YES		88	12	0	
		15	NO	91	9		
		YES		93	7		
		20	NO	93	7		
	25	YES		94	6		
		NO		95	5		
		YES		96	4		
		10	NO	91	9	0	
		YES		92	8		
7	15	NO		96	4		
		YES		96	4		
		20	NO	97	3		
		YES		97	3		
		25	NO	98	2		
	25	YES		98	2		
		NO		99	1		
		YES		99	1		
		10	NO	93	8		
		YES		95	5		
8	15	NO		97	3		
		YES		97	3		
		20	NO	97	3		
		YES		98	2		
		25	NO	99	1		
	25	YES		99	1		
		NO		100			
		YES		100			
		25	NO	100			
		YES		100			
MWX	2	10	NO				100
		YES					100
		25	NO				100
		YES					100
		10	NO		3	97	
	3	YES			14	86	
		25	NO		3	97	
		YES			20	81	
		10	NO	56	38	7	0
		YES		64	34	2	0
4	25	NO		72	27	2	
		YES		79	22	0	
		10	NO	76	23	1	
		YES		78	22	0	
		25	NO	90	10	0	
	5	YES		91	10		
		10	NO	85	15	0	
		NO		95	6		
		YES		96	5		
		25	NO	98	3		
6	25	YES		98	2		
		10	NO	99	2		
		NO		99	1		
		YES		99	1		
		25	NO	98	3		
	7	YES		98	2		
		25	NO	99	2		
		YES		99	1		
		25	NO	99	2		
		YES		99	1		
WILLIAMS 2	10	NO		34	52	14	1
		YES		30	51	18	2
		15	NO	42	49	9	0
		YES		38	47	14	0
		20	YES	42	48	9	0
	25	NO		50	45	6	0
		YES		43	48	8	0
		10	NO				
		YES					
		15	NO				
		YES					
		20	NO				
		YES					
		25	NO				
		YES					

**Table 2**  
**Comparison of Allocation Rules for 4 Dose Experiments**  
POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND  
RELATIVE TO LOG(DOSE)  
MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST	REPS	SAMPLESIZE	SQRTRULE	NOAEL			
				0	1	2	3
3	10	NO		59	40	2	
			YES	59	40	1	
		15	NO	68	32	1	
			YES	68	32	1	
	20	NO		71	29	0	
			YES	70	30	1	
		25	NO	75	25	0	
			YES	74	26	0	
		25	NO	72	29	0	
			YES	76	25	0	
4	15	NO		79	21	0	
			YES	82	18	0	
		20	NO	84	16		
			YES	86	14		
	25	NO		87	13		
			YES	87	13		
5	10	NO		83	17		
			YES	86	14		
		15	NO	88	12		
			YES	90	10		
	20	NO		92	8		
			YES	93	7		
		25	NO	94	6		
			YES	95	5		
6	15	NO		89	11		
			YES	91	9		
		20	NO	94	6		
			YES	95	5		
	25	NO		95	5		
			YES	97	3		
7	20	NO		97	3		
			YES	97	3		
		25	NO	98	2		
			YES	98	2		
	25	NO		99	1		
			YES	99	1		
8	10	NO		96	4		
			YES	97	4		
		15	NO	99	1		
			YES	98	2		
	20	NO		99	1		
			YES	99	1		
		25	NO	99	1		
			YES	100	1		

**Table 3 Effect of Number of Replicates on Power**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST	SQRTRULE	REPS	SAMP SIZE	NOAEL		
				0	1	2
DUNN	NO	2	10		74	26
			15		83	17
			25		87	14
		3	10	23	77	1
			15	20	80	0
			20	20	80	
			25	21	79	0
	4	10		70	31	
			15	77	23	
		20		81	19	
			25	84	16	
		5	10	92	8	
			15	97	3	
			20	98	2	
			25	99	1	
	6	10		0	98	2
			15	0	100	0
			20	0	100	0
			25	0	100	0
		7	10	1	99	0
			15	0	100	0
			20	0	100	0
			25	0	100	0
	8	10		2	98	0
			15	2	98	
			20	1	99	
			25	1	99	
		YES	10		100	
			15		100	
			20		100	
			25		100	
	3	10		17	82	2
			15	16	83	1
			20	14	85	1
			25	13	87	0
		4	10	0	84	16
			15	0	87	13
			20		91	9
			25		93	7
	5	10		2	96	3
			15	1	98	1
			20	1	99	1
			25	0	99	0
		6	10	4	96	0
			15	3	97	0
			20	2	98	0
			25	2	98	0
	7	10		17	83	0
			15	17	83	
			20	16	84	
			25	16	84	
		8	10	26	74	
			15	30	70	
			20	35	65	
			25	37	64	

**Table 3 Effect of Number of Replicates on Power**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST DUNNETT	SQRTRULE NO	REPS 2	SAMP SIZE	NOAEL			
				0	1	2	3
3	2	10	21	52	25	4	
		15	26	54	19	1	
		25	33	54	12	1	
		10	43	53	5	0	
	3	15	53	46	2		
		20	55	44	1		
		25	60	40	0		
		10	59	42	1		
	4	15	67	33	0		
		20	73	27	0		
		25	77	23	0		
		10	70	30	0		
5	2	15	79	21			
		20	85	15			
		25	87	13			
		10	81	19			
	3	15	88	12			
		20	91	9			
		25	93	7			
		10	87	13			
	4	15	93	7			
		20	95	5			
		25	96	4			
		10	92	8			
7	2	15	96	4			
		20	97	3			
		25	98	2			
		10	12	46	34	9	
	3	15	19	49	27	4	
		20	25	51	22	2	
		25	25	51	21	3	
		10	41	55	5		
	4	15	49	49	2		
		20	52	46	2		
		25	57	42	1		
		10	60	40	1		
8	2	15	69	31	0		
		20	74	26	0		
		25	76	24			
		10	74	26	0		
	3	15	81	19	0		
		20	85	15	0		
		25	89	11			
		10	82	18			
	4	15	90	10			
		20	93	7			
		25	94	6			
		10	87	13			
6	2	15	93	7			
		20	95	5			
		25	97	3			
		10	92	8			
	3	15	96	4			
		20	98	2			
		25	98	2			
		10	98	2			

**Table 3 Effect of Number of Replicates on Power**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

NOAEL

0 1 2 3

TEST JONCKH	SQRTRULE NO	REPS 2	SAMP SIZE	75	25	1
3	10	15	82	17	0	
		25	87	14	0	
		15	56	43	2	
		20	61	38	1	
		25	63	36	1	
	15	25	68	32	0	
		10	72	28	0	
		20	78	22	0	
		25	82	18	0	
		25	85	15		
4	10	10	79	22		
		15	85	15		
		20	89	11		
		25	91	9		
		15	86	14		
	15	20	91	9		
		25	93	7		
		25	95	5		
		10	91	9		
		15	96	4		
5	15	20	97	3		
		25	98	2		
		10	94	6		
		15	98	2		
		20	98	2		
	20	25	99	1		
		10	86	12	3	
		15	90	8	2	
		20	91	7	2	
		25	92	7	1	
6	10	10	66	31	4	
		15	73	25	2	
		20	76	22	2	
		25	78	20	2	
	15	10	75	25	1	
		15	81	19	0	
		20	86	15		
		25	86	14		
		10	85	16	0	
7	15	15	90	10		
		20	92	8		
		25	93	7		
		10	91	9		
	20	15	95	5		
		20	96	4		
		25	97	3		
		10	92	8		
		15	96	4		
8	20	20	97	3		
		25	98	2		
		10	96	4		
		15	98	2		
		20	99	1		
	25	25	99	1		

**Table 3 Effect of Number of Replicates on Power**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST JONCKHX	SQRTRULE NO	REPS 2	SAMPSIZE	NOAEL			
				0	1	2	3
YES	2	10		75	25	1	
		25		88	13	0	
		3	10	28	68	4	
		25		34	66	1	
		4	10	57	43	0	
		25		73	28		
		5	10	76	24		
		25		90	10		
	3	10		86	15		
		25		95	6		
		7	25	98	3		
		8	25	99	2		
		2	10	62	31	7	
		25		73	24	4	
		3	10	49	48	3	
		25		62	37	2	
MW	2	10		65	35	1	
		25		79	22	0	
		5	10	78	22	0	
		25		90	10		
		6	25	96	5		
		7	25	98	2		
		8	25	99	1		
		2	10		100		
	3	15			100		
		25			100		
		10			100		
		15			100		
		20			100		
		25			100		
		4	10	55	38	7	0
		15		64	32	4	0
	5	20		68	30	2	
		25		72	27	2	
		10		78	21	1	
		15		85	15	0	
		20		89	11	0	
		25		91	9	0	
		6	10	86	14	0	
		15		91	9		
	7	20		93	7		
		25		95	5		
		10		91	9	0	
		15		96	4		
		20		97	3		
		25		98	2		
		8	10	93	8		
		15		97	3		
	25	20		97	3		
		25		99	1		

**Table 3 Effect of Number of Replicates on Power**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

NOAEL

0 1 2 3

TEST	SQRTRULE	REPS	SAMP SIZE				
YES		2	10				100
			15				100
			20				100
			25				100
		3	10				100
			15				100
			20				100
			25				100
		4	10	64	34	2	0
			15	71	27	1	0
			20	76	23	1	
			25	78	22	0	0
		5	10	78	22	1	
			15	84	15	0	
			20	87	13	0	
			25	90	10	0	
		6	10	88	12	0	
			15	93	7		
			20	94	6		
			25	96	4		
		7	10	92	8		
			15	96	4		
			20	97	3		
			25	98	2		
		8	10	95	5		
			15	97	3		
			20	98	2		
			25	99	1		
MWX	NO	2	10				100
			25				100
		3	10			3	97
			25			3	97
		4	10	56	38	7	0
			25	72	27	2	
		5	10	76	23	1	
			25	90	10	0	
		6	10	85	15	0	
			25	95	6		
		7	25	98	3		
		8	25	99	2		
	YES	2	10				100
			25				100
		3	10			14	86
			25			20	81
		4	10	64	34	2	0
			25	79	22	0	
		5	10	78	22	0	
			25	91	10		
		6	25	96	5		
		7	25	98	2		
		8	25	99	1		

**Table 3 Effect of Number of Replicates on Power**

POWER SIMULATIONS FOR 4 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.2918 0.5836 0.8754

TEST	SQRTRULE	REPS	SAMPSIZE	NOAEL			
				0	1	2	3
WILLIAMS NO	2	10		34	52	14	1
		15		42	49	9	0
		25		50	45	6	0
		10		59	40	2	
		15		68	32	1	
	3	20		71	29	0	
		25		75	25	0	
		10		72	29	0	
		15		79	21	0	
	4	20		84	16		
		25		87	13		
		10		83	17		
	5	15		88	12		
		20		92	8		
		25		94	6		
		10		89	11		
	6	15		94	6		
		20		95	5		
		25		97	3		
		10		93	7		
	7	15		97	3		
		20		98	2		
		25		99	1		
		10		96	4		
		15		99	1		
	8	20		99	1		
		25		99	1		
		10		98	2		
		15		99	1		
YES	2	10		30	51	18	2
		15		38	47	14	0
		20		42	48	9	0
		25		43	48	8	0
	3	10		59	40	1	
		15		68	32	1	
		20		70	30	1	
		25		74	26	0	
	4	10		76	25	0	
		15		82	18	0	
		20		86	14		
		25		87	13		
	5	10		86	14		
		15		90	10		
		20		93	7		
		25		95	5		
	6	10		91	9		
		15		95	5		
		20		97	3		
		25		97	3		
	7	10		95	6		
		15		97	3		
		20		98	2		
		25		99	1		
		10		97	4		
	8	15		98	2		
		20		99	1		
		25		100	1		

**Table 4 Comparison of Tests for 5 Dose Experiments**

POWER SIMULATIONS FOR 5 DOSES, LINEAR DOSE-RESPONSE TREND  
 RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, DOSES=0 0.21885 0.4377 0.65655 0.8754

REPS	SAMP SIZE	SQRTRULE TEST	NOAEL				
			0	1	2	3	4
2	10	DUNN		6	69	26	
		DUNNETT	11	35	37	15	3
		JONCKH		55	41	5	0
		JONCKHX		55	39	6	0
		MW				100	
		MWX				100	
		WILLIAMS	25	43	26	6	1
	YES	DUNN				100	
		DUNNETT	5	26	40	23	7
		JONCKH		65	22	13	1
		JONCKHX		39	47	13	1
		MW				100	
		MWX				100	
		WILLIAMS	21	43	30	7	1
		DUNN		6	76	19	
15	NO	DUNNETT	14	42	33	9	1
		JONCKH		61	37	2	0
		MW				100	
		WILLIAMS	29	47	20	3	0
		DUNN				100	
		DUNNETT	8	34	38	16	3
		JONCKH		71	19	9	1
	YES	MW				100	
		WILLIAMS	25	45	26	4	0
		DUNN				100	
		DUNNETT	10	37	37	14	2
		JONCKH		73	18	8	1
		MW				100	
		WILLIAMS	27	48	22	3	0
		DUNN		6	82	13	
20	YES	DUNNETT	18	47	31	6	1
		JONCKH		68	31	1	0
		JONCKHX		68	30	2	
		MW				100	
		MWX				100	
		WILLIAMS	34	49	18	1	0
		DUNN				100	
	NO	DUNNETT	13	39	37	11	2
		JONCKH		76	17	7	1
		JONCKHX		50	42	8	0
		MW				100	
		MWX				100	
		WILLIAMS	31	48	19	3	0
		DUNN		0	56	43	2
		DUNNETT	24	54	21	2	0
3	10	JONCKH	37	51	12	0	
		JONCKHX		19	62	19	1
		MW				100	
		MWX				1	99
		WILLIAMS	42	49	10	0	
		DUNN		0	49	49	3
		DUNNETT	20	56	24	2	0
	NO	JONCKH	43	47	10	1	
		JONCKHX		28	56	16	1
		MW				100	
		MWX				6	94
		WILLIAMS	41	51	9	0	0

**Table 4 Comparison of Tests for 5 Dose Experiments**

POWER SIMULATIONS FOR 5 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, DOSES=0 0.21885 0.4377 0.65655 0.8754

REPS	SAMP SIZE	SQR RULE TEST	NOAEL				
			0	1	2	3	4
15	NO	DUNN		0	63	36	1
		DUNNETT	30	56	14	0	
		JONCKH	42	51	6	0	
		MW				100	
		WILLIAMS	49	46	6	0	
	YES	DUNN		0	54	44	1
		DUNNETT	27	57	15	1	
		JONCKH	50	44	6	0	
		MW				100	
		WILLIAMS	49	47	4	0	
20	NO	DUNN		0	67	33	0
		DUNNETT	31	57	11	0	
		JONCKH	44	51	4		
		MW				100	
		WILLIAMS	50	46	4	0	
	YES	DUNN		0	59	41	1
		DUNNETT	29	59	12	0	
		JONCKH	53	43	4	0	
		MW				100	
		WILLIAMS	52	45	3	0	
25	NO	DUNN		0	69	32	1
		DUNNETT	35	57	9	0	
		JONCKH	48	49	4	0	
		JONCKHX	24	68	8		
		MW				100	
		MWX				0	100
	YES	WILLIAMS	54	43	3	0	
		DUNN			61	39	1
		DUNNETT	34	57	10	0	
		JONCKH	56	41	4	0	
		JONCKHX	40	54	7		
4	10	NO	MW			100	
			MWX			9	91
			WILLIAMS	56	43	2	
			DUNN		4	87	10
			DUNNETT	34	57	9	0
			JONCKH	52	44	6	0
			JONCKHX	37	58	5	
			MW			100	
			MWX	7	11	20	29
			WILLIAMS	53	44	3	
15	NO	YES	DUNN		14	81	6
			DUNNETT	34	58	8	0
			JONCKH	50	47	3	
			JONCKHX	50	46	5	
			MW	45	37	17	1
			MWX	47	41	11	0
			WILLIAMS	57	42	2	
			DUNN		3	90	7
			DUNNETT	41	54	5	
			JONCKH	57	40	3	
	YES	MW				100	
		WILLIAMS	59	39	2		
		DUNN			14	83	3
		DUNNETT	41	55	4		
		JONCKH	56	43	1		
	MW		53	37	10	0	
		WILLIAMS	62	37	1		

**Table 4 Comparison of Tests for 5 Dose Experiments**

POWER SIMULATIONS FOR 5 DOSES, LINEAR DOSE-RESPONSE TREND  
RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, DOSES=0 0.21885 0.4377 0.65655 0.8754

REPS	SAMP SIZE	SQRTRULE	TEST	NOAEL			
				0	1	2	3
20	NO	DUNN			3	93	4
		DUNNETT		45	52	3	
		JONCKH		63	35	2	
		MW					100
		WILLIAMS		65	35	1	
	YES	DUNN			13	85	2
		DUNNETT		45	53	3	
		JONCKH		60	39	1	
		MW		57	35	7	0
		WILLIAMS		67	32	1	
25	NO	DUNN			3	93	4
		DUNNETT		48	50	2	
		JONCKH		65	34	1	
		JONCKHX		47	51	1	
		MW					100
	YES	MWX		10	15	23	24
		WILLIAMS		66	34	1	
		DUNN			14	85	2
		DUNNETT		49	49	2	
		JONCKH		63	36	1	
5	10	JONCKHX		63	36	1	
		MW		61	34	6	0
		MWX		62	35	3	
		WILLIAMS		70	30	0	
		DUNN			16	83	2
	YES	DUNNETT		43	54	4	0
		JONCKH		57	42	2	
		JONCKHX		56	43	2	
		MW		56	36	9	1
		MWX		55	37	8	0
15	NO	WILLIAMS		63	37	1	
		DUNN			41	59	0
		DUNNETT		46	51	3	
		JONCKH		65	34	1	
		MW		56	39	5	0
	YES	WILLIAMS		67	33	0	
		DUNN			16	83	1
		DUNNETT		51	48	1	
		JONCKH		65	35	1	
		MW		64	31	5	0
20	NO	WILLIAMS		69	30	0	
		DUNN			46	54	0
		DUNNETT		54	46	1	
		JONCKH		72	28	0	
		MW		64	34	2	0
	YES	WILLIAMS		74	26	0	
		DUNN			15	84	0
		DUNNETT		57	43	1	
		JONCKH		69	30	0	
		MW		68	28	3	0

**Table 4 Comparison of Tests for 5 Dose Experiments**

POWER SIMULATIONS FOR 5 DOSES, LINEAR DOSE-RESPONSE TREND  
RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, DOSES=0 0.21885 0.4377 0.65655 0.8754

REPS	SAMP SIZE	SQR RULE	TEST	NOAEL				
				0	1	2	3	4
25	NO	DUNN	DUNN	15	86	0		
			DUNNETT	60	40	1		
			JONCKH	73	27	0		
			JONCKHX	72	28	0		
			MW	73	25	3	0	
	YES	WILLIAMS	MWX	72	26	3		
			WILLIAMS	78	22	0		
			DUNN	52	49	0		
			DUNNETT	64	36	0		
			JONCKH	80	20	0		
6	10	NO	JONCKHX	74	26			
			MW	73	27	1		
			MWX	74	26	1		
			WILLIAMS	82	18	0		
			DUNN	36	63	0		
	YES	WILLIAMS	DUNNETT	51	48	1		
			JONCKH	63	36	1		
			MW	63	34	3	0	
			WILLIAMS	69	31	0		
			DUNN	0	64	36	0	
15	NO	WILLIAMS	DUNNETT	55	44	1		
			JONCKH	71	29	0		
			MW	70	29	1		
			WILLIAMS	74	26	0		
			DUNN	40	60	0		
	YES	WILLIAMS	DUNNETT	61	39	0		
			JONCKH	72	28	0		
			MW	72	27	1		
			WILLIAMS	77	23	0		
			DUNN	72	28	0		
20	NO	WILLIAMS	DUNNETT	64	36	0		
			JONCKH	78	22			
			MW	78	21	0		
			WILLIAMS	81	19			
			DUNN	42	58	0		
	YES	WILLIAMS	DUNNETT	66	34	0		
			JONCKH	77	23	0		
			MW	77	23	1		
			WILLIAMS	82	18	0		
			DUNN	0	76	24		
25	NO	WILLIAMS	DUNNETT	69	31	0		
			JONCKH	82	18	0		
			MW	82	18	0		
			WILLIAMS	84	16			
			DUNN	44	57			
	YES	WILLIAMS	DUNNETT	70	30	0		
			JONCKH	80	21	0		
			JONCKHX	79	21	0		
			MW	80	20	0		
			MWX	79	21	0		
			WILLIAMS	84	17			
			DUNN	0	80	21		
			DUNNETT	74	27	0		
			JONCKH	85	15	0		
			JONCKHX	83	17			
			MW	85	15	0		
			MWX	84	16			
			WILLIAMS	87	13			

**Table 4 Comparison of Tests for 5 Dose Experiments**

POWER SIMULATIONS FOR 5 DOSES, LINEAR DOSE-RESPONSE TREND

RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, DOSES=0 0.21885 0.4377 0.65655 0.8754

REPS	SAMP SIZE	SQR RULE	TEST	NOAEL				
				0	1	2	3	4
7	10	NO	DUNN	0	60	40		
			DUNNETT	59	40	1		
			JONCKH	72	27	0		
			MW	72	27	1		
			WILLIAMS	76	24	0		
	15	NO	DUNN		66	34		
			DUNNETT	67	33	0		
			JONCKH	79	21	0		
			MW	79	21	0		
			WILLIAMS	82	18	0		
20	NO	YES	DUNN	1	91	8		
			DUNNETT	69	31	0		
			JONCKH	83	17	0		
			MW	83	17	0		
			WILLIAMS	85	15			
	YES	NO	DUNN		71	29		
			DUNNETT	73	27			
			JONCKH	83	17			
			MW	83	17	0		
			WILLIAMS	86	14			
25	NO	YES	DUNN	1	94	5		
			DUNNETT	75	25	0		
			JONCKH	87	13			
			MW	87	13	0		
			WILLIAMS	89	11			
	YES	NO	DUNN	0	74	27		
			DUNNETT	77	24	0		
			JONCKH	86	14			
			JONCKHX	84	16			
			MW	86	14	0		
8	15	NO	MWX	85	15	0		
			WILLIAMS	89	11			
			DUNN	1	95	4		
			DUNNETT	79	21			
			JONCKH	90	11	0		
	YES	YES	JONCKHX	88	12			
			MW	90	11			
			MWX	88	12			
			WILLIAMS	91	9			
			DUNN	3	94	3		
20	NO	NO	DUNNETT	76	24	0		
			JONCKH	86	14			
			MW	83	17	0		
			WILLIAMS	88	12			
			DUNN	3	94	3		
	YES	YES	DUNNETT	77	23			
			JONCKH	89	11			
			MW	89	11			
			WILLIAMS	90	10			
			DUNN		87	13		
8	15	NO	DUNNETT	80	20			
			JONCKH	89	11			
			MW	87	13			
	YES	YES	WILLIAMS	91	9			
			DUNN	3	95	2		
			DUNNETT	82	18			

**Table 4 Comparison of Tests for 5 Dose Experiments**

POWER SIMULATIONS FOR 5 DOSES, LINEAR DOSE-RESPONSE TREND  
RELATIVE TO LOG(DOSE)  
MAX EFFECT=0.8754, DOSES=0 0.21885 0.4377 0.65655 0.8754

REPS	SAMP SIZE	SQRTRULE	TEST	NOAEL			
				0	1	2	3
8	20	YES	WILLIAMS	92	8		
			DUNN		90	11	
		NO	DUNNETT	84	17		
			JONCKH	91	9		
			JONCKHX	89	11		
			MW	89	11	0	
			MWX	89	11		
	25	YES	WILLIAMS	93	8		
			DUNN	2	97	1	
			DUNNETT	85	16	0	
			JONCKH	92	8		
			JONCKHX	93	7		
			MW	92	8		
			MWX	93	7		
			WILLIAMS	94	6		

**Table 5 Comparison of Allocation Rules for 5 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST DUNN	REPS	SAMP SIZE	SQR RULE	NOAEL				
				0	1	2	3	4
2	10	NO			6	69	26	
			YES				100	
		15	NO		6	76	19	
			YES				100	
			YES				100	
	25	NO			6	81	13	
			YES				100	
		20	NO		0	56	43	2
			YES		0	49	49	3
			NO		0	63	36	1
3	15	YES			0	54	44	1
			NO		0	67	33	0
		20	YES		0	59	41	1
			NO		0	69	31	0
			YES		0	61	38	1
	25	NO			4	87	10	0
			YES		14	81	6	0
		20	NO		3	90	7	
			YES		14	83	3	
			NO		3	93	4	
4	10	YES			13	85	2	
			NO		3	93	3	
		15	NO		3	93	3	
			YES		14	85	1	
			NO		16	83	2	
	25	YES			41	59	0	
			NO		16	83	1	
		20	YES		46	54	0	
			NO		15	84	0	
			YES		50	50	0	
5	15	NO			14	86	0	
			YES		51	49	0	
		20	NO		37	63	0	
			YES		0	65	36	0
			NO		40	60	0	
	25	YES			72	28	0	
			NO		42	58	0	
		20	YES		0	76	24	
			NO		43	57		
			YES		0	80	20	
6	10	NO			0	60	40	
			YES		3	83	14	
		15	NO		66	34		
			YES		1	91	8	
			NO		71	29		
	25	YES			1	94	5	
			NO		0	73	27	
		20	YES		1	95	4	
			NO		75	25		
			YES		4	89	7	
7	15	NO			84	16		
			YES		3	94	3	
		20	NO		87	13		
			YES		3	95	2	
			NO		89	11		
	25	YES			2	97	1	
			NO					

**Table 5 Comparison of Allocation Rules for 5 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST	REPS	SAMP SIZE	SQR RULE	NOAEL				
				0	1	2	3	4
DUNNETT	2	10	NO	11	35	37	15	3
			YES	5	26	40	23	7
		15	NO	14	42	33	9	1
			YES	8	34	38	16	3
		20	YES	11	37	37	14	2
	3	25	NO	18	46	30	6	0
			YES	12	39	36	11	1
		10	NO	24	54	21	2	0
			YES	20	56	24	2	0
		15	NO	30	56	14	0	
4	20		YES	27	57	15	1	
		25	NO	31	57	11	0	
			YES	29	59	12	0	
		25	NO	35	56	9	0	
			YES	33	57	10	0	
	25	10	NO	34	57	9	0	
			YES	34	58	8	0	
		15	NO	41	54	5		
			YES	41	55	4		
		20	NO	45	52	3		
5	25		YES	45	53	3		
		25	NO	48	49	2		
			YES	49	49	2	0	
		10	NO	43	54	4	0	
			YES	45	53	3		
	30	15	NO	51	48	1		
			YES	54	46	1		
		20	NO	57	43	1		
			YES	58	42	0		
		25	NO	60	39	0		
6	30		YES	64	36	0		
		10	NO	52	47	1		
			YES	55	45	1		
		15	NO	61	39	0		
			YES	64	36	0		
	35	20	NO	66	34	0		
			YES	69	31	0		
		25	NO	70	30	0		
			YES	73	27	0		
		30	NO	59	41	1		
7	35		YES	61	39	0		
		10	NO	67	33	0		
			YES	69	31	0		
		15	NO	73	27			
			YES	75	25	0		
	40	20	NO	76	24	0		
			YES	79	21			
		25	NO	66	34	0		
			YES	69	31	0		
		30	NO	76	24	0		
8	40		YES	77	23			
		10	NO	80	20			
			YES	82	18			
		15	NO	83	17			
			YES	85	15	0		

**Table 5 Comparison of Allocation Rules for 5 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST	REPS	SAMP SIZE	SQR RULE	NOAEL				
				0	1	2	3	4
JONCKH	2	10	NO		55	41	5	0
			YES		65	22	13	1
		15	NO		61	37	2	0
			YES		71	19	9	1
		20	YES		74	18	8	1
	3	25	NO		68	31	1	0
			YES		76	17	7	0
		10	NO	37	51	12	0	
			YES	43	47	10	1	
		15	NO	42	51	6	0	
4	4		YES	50	44	6	0	
		20	NO	44	51	4		
			YES	53	43	4	0	
		25	NO	48	48	4	0	
			YES	56	41	4	0	
	5	10	NO	52	44	6	0	
			YES	50	47	3		
		15	NO	57	40	3		
			YES	56	43	1		
		20	NO	63	35	2		
6	6		YES	60	39	1		
		25	NO	65	34	1		
			YES	63	36	0		
		10	NO	57	42	2		
			YES	65	35	1		
	7	15	NO	65	35	1		
			YES	72	28	0		
		20	NO	69	30	0		
			YES	76	24	0		
		25	NO	73	27	0		
8	8		YES	80	20	0		
		10	NO	66	34	1		
			YES	71	29	0		
		15	NO	72	28	0		
			YES	78	22			
	9	20	NO	77	23	0		
			YES	82	18	0		
		25	NO	79	21	0		
			YES	85	15	0		
		10	NO	72	27	0		
10	10		YES	77	23	0		
		15	NO	79	21	0		
			YES	83	17	0		
		20	NO	83	17			
			YES	87	13			
	11	25	NO	86	14	0		
			YES	89	11	0		
		10	NO	89	11			
			YES	91	9			
		15	NO	91	9			
11	11		YES	92	8			

**Table 5 Comparison of Allocation Rules for 5 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST	REPS	SAMP SIZE	SQR RULE	NOAEL				
				0	1	2	3	4
JONCKHX	2	10	NO		55	39	6	0
			YES		39	47	13	1
		25	NO		68	30	2	
			YES		50	42	8	0
			NO	19	62	19	1	
	3	10	YES	28	56	16	1	
			NO	24	68	8		
		25	YES	40	54	7		
			NO	37	58	5		
			YES	50	46	5		
5	4	10	NO	47	51	1		
			YES	63	36	1		
		25	NO	56	43	2		
			YES	58	41	1		
			NO	72	28	0		
	5	10	YES	74	26			
			NO	68	31	1		
		25	YES	68	32	0		
			NO	79	21	0		
			YES	83	17			
6	6	10	NO	69	31	0		
			YES	76	24	0		
		25	NO	84	16			
			YES	88	12			
			NO	77	23	0		
	7	10	YES	82	18			
			NO	89	11			
		25	YES	93	7			
			NO					
			YES					
MW	2	10	NO			100		
			YES			100		
		15	NO			100		
			YES			100		
			NO			100		
	3	10	YES			100		
			NO			100		
		15	YES			100		
			NO			100		
			YES			100		
4	4	10	NO			100		
			YES	45	37	17	1	0
		15	NO			100		
			YES	53	37	10	0	
			NO			100		
	5	20	YES	57	35	7	0	
			NO			100		
		25	YES	61	33	6	0	
			NO	56	36	9	1	0
			YES	56	40	5	0	
5	5	10	NO	64	31	5	0	
			YES	64	34	2	0	
		15	NO	68	28	3	0	
			YES	69	29	1		
			NO	72	25	2	0	
	6	25	YES	73	26	1		
			NO					

**Table 5 Comparison of Allocation Rules for 5 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST MW	REPS 6	SAMP SIZE	SOR RULE	NOAEL				
				0	1	2	3	4
7	10	10	NO	66	32	3	0	
			YES	71	29	1		
		15	NO	72	27	1		
			YES	78	21	0		
			NO	77	23	1		
	25	20	YES	82	18	0		
			NO	79	20	0		
		25	YES	85	15	0		
			NO	79	20	0		
			YES	83	17	0		
8	10	10	NO	72	27	1		
			YES	77	23	1		
		15	NO	79	21	0		
			YES	83	17	0		
			NO	83	17	0		
	25	20	YES	87	13	0		
			NO	86	14	0		
		25	YES	89	11	0		
			NO	77	23	0		
			YES	83	17	0		
MWX	2	10	NO				100	
			YES				100	
		25	NO				100	
			YES				100	
			NO				1	99
	3	10	YES				6	94
			NO				0	100
		25	NO				9	91
			YES				7	11
			NO				47	41
4	10	10	YES	20	29	33		
			NO	11	11	0		
		25	YES	10	15	23	24	27
			NO	62	35	3		
			YES	62	35	3		
	25	10	NO	55	37	8	0	
			YES	56	41	4		
		25	NO	72	26	3		
			YES	74	26	1		
			NO	68	30	2		
5	10	10	YES	68	31	1		
			NO	79	21	0		
		25	YES	84	16			
			NO	69	30	1		
			YES	76	23	1		
	25	10	NO	85	15	0		
			YES	88	12			
		25	NO	77	23	0		
			YES	83	17			
			NO	89	11			
6	10	10	YES	93	7			
			NO					
		25	NO					
			YES					
			NO					
	25	10	NO					
			YES					
		25	NO					
			YES					
			NO					
7	10	10	NO					
			YES					
		25	NO					
			YES					
			NO					
	25	10	NO					
			YES					
		25	NO					
			YES					
			NO					
8	10	10	NO					
			YES					
		25	NO					
			YES					
			NO					
	25	10	NO					
			YES					
		25	NO					
			YES					
			NO					

**Table 5 Comparison of Allocation Rules for 5 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST	REPS	SAMP SIZE	SQR RULE	NOAEL				
				0	1	2	3	4
WILLIAMS 2	10	NO		25	43	26	6	1
		YES		21	43	30	7	1
		15	NO	29	47	20	3	0
			YES	25	45	26	4	0
		20	YES	28	48	22	3	0
	25	NO		34	49	17	1	0
			YES	31	48	19	2	0
		3	NO	42	49	10	0	
			YES	41	51	9	0	0
		15	NO	49	46	6	0	
			YES	49	47	4	0	
WILLIAMS 3	20	NO		50	46	4	0	
			YES	52	45	3	0	
		25	NO	54	43	3	0	
			YES	55	42	2		
		4	NO	53	44	3		
	25		YES	57	42	2		
		15	NO	59	39	2		
			YES	62	37	1		
		20	NO	65	35	1		
			YES	67	32	1		
5	25	NO		66	33	0		
			YES	70	29	0		
		5	NO	63	37	1		
			YES	67	33	0		
		15	NO	69	30	0		
	10		YES	74	26	0		
		20	NO	74	25	0		
			YES	77	23	0		
		25	NO	78	22	0		
			YES	82	18	0		
6	10	NO		72	29	0		
			YES	74	26	0		
		15	NO	77	23	0		
			YES	81	19			
		20	NO	82	18	0		
	25		YES	84	16			
		25	NO	84	16			
			YES	87	13			
		7	NO	75	25	0		
			YES	80	20			
8	15	NO		82	18	0		
			YES	85	15			
		20	NO	86	14			
			YES	89	11			
		25	NO	89	11			
	10		YES	91	9			
		20	NO	82	18			
			YES	84	16			
		15	NO	88	12			
			YES	90	10			
25	10	NO		91	9			
			YES	92	8			
	15	NO		93	7			
			YES	94	6			

**Table 6 Effect of Number of Replicates on Power**

POWER COMPARISON OF REPLICATE COUNT FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST DUNN	SQRTRULE NO	REPS 2	SAMP SIZE	NOAEL				
				0	1	2	3	4
3	10	10			6	69	26	
		15			6	76	19	
		25			6	81	13	
		0	56	43	2			
	15	10			0	63	36	1
		15			0	67	33	0
		20			0	69	31	0
		25			4	87	10	0
	20	10			3	90	7	
		15			3	93	4	
		25			3	93	3	
		0	83	2				
4	15	10			16	83	1	
		15			15	84	0	
		20			14	86	0	
		25			37	63	0	
	20	10			40	60	0	
		15			42	58	0	
		25			43	57		
		0	60	40				
	25	10			66	34		
		15			71	29		
		20			0	73	27	
		25			75	25		
5	10	10			84	16		
		15			87	13		
		20			89	11		
		0	83	2				
	15	10			14	81	6	0
		15			14	83	3	
		20			13	85	2	
		25			14	85	1	
	20	10			41	59	0	
		15			46	54	0	
		20			50	50	0	
		25			51	49	0	
6	15	10			0	65	36	0
		15			72	28	0	
		20			0	76	24	
		25			0	80	20	
	20	10			3	83	14	
		15			1	91	8	
		20			1	94	5	
		25			1	95	4	
	25	10			4	89	7	
		15			3	94	3	
		20			3	95	2	
		25			2	97	1	
YES	2	10				100		
		15				100		
		20				100		
		25				100		
	3	10			0	49	49	3
		15			0	54	44	1
		20			0	59	41	1
		25			0	61	38	1
	4	10			14	81	6	0
		15			14	83	3	
		20			13	85	2	
		25			14	85	1	
8	5	10			41	59	0	
		15			46	54	0	
		20			50	50	0	
		25			51	49	0	
	6	10			0	65	36	0
		15			72	28	0	
		20			0	76	24	
		25			0	80	20	
	7	10			3	83	14	
		15			1	91	8	
		20			1	94	5	
		25			1	95	4	
8	10	10			4	89	7	
		15			3	94	3	
		20			3	95	2	
		25			2	97	1	

**Table 6 Effect of Number of Replicates on Power**

POWER COMPARISON OF REPLICATE COUNT FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST	DUNNETT	SQRT RULE	REPS	SAMP SIZE	NOAEL				
					0	1	2	3	4
NO	TEST DUNNETT	NO	2	10	11	35	37	15	3
				15	14	42	33	9	1
				25	18	46	30	6	0
				3	10	24	54	21	0
				15	30	56	14	0	
			4	20	31	57	11	0	
				25	35	56	9	0	
				4	10	34	57	9	0
				15	41	54	5		
				20	45	52	3		
YES	TEST DUNNETT	YES	2	25	48	49	2		
				5	10	43	54	4	0
				15	51	48	1		
				20	57	43	1		
				25	60	39	0		
			3	6	10	52	47	1	
				15	61	39	0		
				20	66	34	0		
			4	25	70	30	0		
				7	10	59	41	1	
				15	67	33	0		
				20	73	27			
				25	76	24	0		
NO	TEST DUNNETT	NO	2	8	10	66	34	0	
				15	76	24	0		
				20	80	20			
				25	83	17			
				3	10	5	26	23	7
				15	8	34	38	16	3
				20	11	37	37	14	2
				25	12	39	36	11	1
			4	4	10	20	56	24	0
				15	27	57	15	1	
				20	29	59	12	0	
				25	33	57	10	0	
				5	10	34	58	8	0
YES	TEST DUNNETT	YES	2	15	15	41	55	4	
				20	45	53	3		
				25	49	49	2	0	
				6	10	45	53	3	
				15	54	46	1		
			3	20	58	42	0		
				25	64	36	0		
				7	10	55	45	1	
				15	64	36	0		
				20	69	31	0		
NO	TEST DUNNETT	NO	2	25	73	27	0		
				10	61	39	0		
				15	69	31	0		
				20	75	25	0		
				25	79	21			
			3	8	10	69	31	0	
				15	77	23			
				20	82	18			
				25	85	15	0		

**Table 6 Effect of Number of Replicates on Power**

POWER COMPARISON OF REPLICATE COUNT FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST JONCKH	SQRTRULE NO	REPS 2	SAMPSIZE	NOAEL				
				0	1	2	3	4
3	NO	2	10		55	41	5	0
			15		61	37	2	0
			25		68	31	1	0
			10		37	51	12	0
			15		42	51	6	0
	YES	2	20		44	51	4	0
			25		48	48	4	0
			10		52	44	6	0
			15		57	40	3	
			20		63	35	2	
4	NO	2	25		65	34	1	
			10		57	42	2	
			15		65	35	1	
			20		69	30	0	
			25		73	27	0	
	YES	2	10		66	34	1	
			15		72	28	0	
			20		77	23	0	
			25		79	21	0	
			10		72	27	0	
5	NO	2	15		79	21	0	
			20		83	17		
			25		86	14	0	
			10		80	20	0	
			15		86	14		
	YES	2	20		89	11		
			25		91	9		
			10		65	22	13	1
			15		71	19	9	1
			20		74	18	8	1
6	NO	2	25		76	17	7	0
			10		43	47	10	1
			15		50	44	6	0
			20		53	43	4	0
			25		56	41	4	0
	YES	2	10		50	47	3	
			15		56	43	1	
			20		60	39	1	
			25		63	36	0	
			10		65	35	1	
7	NO	2	15		72	28	0	
			20		76	24	0	
			25		80	20	0	
			10		71	29	0	
			15		78	22		
	YES	2	20		82	18	0	
			25		85	15	0	
			10		77	23	0	
			15		83	17	0	
			20		87	13		
8	NO	2	25		89	11	0	
			10		83	17		
			15		89	11		
			20		91	9		
			25		92	8		

**Table 6 Effect of Number of Replicates on Power**

POWER COMPARISON OF REPLICATE COUNT FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST JONCKHX	SQRTRULE NO	REPS	SAMP SIZE	NOAEL				
				0	1	2	3	4
YES	2	10		55	39	6	0	
			25	68	30	2		
		3	10	19	62	19	1	
			25	24	68	8		
		4	10	37	58	5		
			25	47	51	1		
		5	10	56	43	2		
			25	72	28	0		
	8	10		68	31	1		
			25	79	21	0		
		7	10	69	31	0		
			25	84	16			
		10		77	23	0		
			25	89	11			
		3	10	39	47	13	1	
			25	50	42	8	0	
			10	28	56	16	1	
			25	40	54	7		
			4	10	50	46	5	
			25	63	36	1		
			5	10	58	41	1	
			25	74	26			
MW	2	10		68	32	0		
			25	83	17			
		7	10	76	24	0		
			25	88	12			
		8	10	82	18			
			25	93	7			
		3	10			100		
			15			100		
			25			100		
			10			100		
			15			100		
			20			100		
			25			100		
			4	10		100		
	4	15				100		
			20			100		
		5	10			100		
			15			100		
		6	20			100		
			25			100		
		7	10	56	36	9	1	0
			15	64	31	5	0	
			20	68	28	3	0	
			25	72	25	2	0	
			10	66	32	3	0	
			15	72	27	1		
			20	77	23	1		
			25	79	20	0		
	8	10		72	27	1		
			15	79	21	0		
		20		83	17	0		
			25	86	14	0		
		15	10	77	23	0		
			20	83	17	0		
		25		87	13			
				89	11	0		

**Table 6 Effect of Number of Replicates on Power**

POWER COMPARISON OF REPLICATE COUNT FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST MW	SQRTRULE	REPS	SAMP SIZE	NOAEL				
				0	1	2	3	4
TEST MW	YES	2	10					100
			15					100
			20					100
			25					100
			30	10				100
		3	15					100
			20					100
			25					100
			30	10				100
			35					100
TEST MW	NO	2	10	45	37	17	1	0
			15	53	37	10	0	
			20	57	35	7	0	
			25	61	33	6	0	
			30	56	40	5	0	
		3	35	64	34	2	0	
			40	69	29	1		
			45	73	26	1		
			50	71	29	1		
			55	78	21	0		
TEST MW	NO	4	60	82	18	0		
			65	85	15	0		
			70	77	23	1		
			75	83	17	0		
			80	87	13	0		
		5	85	89	11	0		
			90	83	17			
			95	89	11			
			100	92	8			
			105					
TEST MW	NO	6	10					100
			15					100
			20					100
			25					100
			30	77	23	1		
		7	35	83	17	0		
			40	87	13	0		
			45	89	11	0		
			50	91	9			
			55	92	8			
TEST MW	NO	8	60					
			65					
			70					
			75					
			80					
		9	85					
			90					
			95					
			100					
			105					
TEST MW	NO	10	10					
			15					
			20					
			25					
			30	77	23	1		
		11	35	83	17	0		
			40	87	13	0		
			45	89	11	0		
			50	91	9			
			55	92	8			
TEST MW	NO	12	10					
			15					
			20					
			25					
			30	77	23	1		
		13	35	83	17	0		
			40	87	13	0		
			45	89	11	0		
			50	91	9			
			55	92	8			
TEST MW	NO	14	10					
			15					
			20					
			25					
			30	47	41	11	0	
		15	35	62	35	3		
			40	56	41	4		
			45	74	26	1		
			50	68	31	1		
			55	84	16			
TEST MW	NO	16	10					
			15					
			20					
			25					
			30	76	23	1		
		17	35	88	12			
			40	83	17			
			45	93	7			
			50					
			55					

**Table 6 Effect of Number of Replicates on Power**  
 POWER COMPARISON OF REPLICATE COUNT FOR 5 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.21885 0.4377 0.65655 0.8754

TEST	SQRTRULE	REPS	SAMP SIZE	NOAEL				
				0	1	2	3	4
WILLIAMS NO	2	10		25	43	26	6	1
		15		29	47	20	3	0
		25		34	49	17	1	0
		3	10	42	49	10	0	
		15		49	46	6	0	
		20		50	46	4	0	
		25		54	43	3	0	
		4	10	53	44	3		
		15		59	39	2		
		20		65	35	1		
		25		66	33	0		
		5	10	63	37	1		
		15		69	30	0		
		20		74	25	0		
		25		78	22	0		
		6	10	72	29	0		
		15		77	23	0		
		20		82	18	0		
		25		84	16			
YES	2	10		75	25	0		
		15		82	18	0		
		20		86	14			
		25		89	11			
		8	10	82	18			
		15		88	12			
		20		91	9			
		25		93	7			
		3	10	21	43	30	7	1
		15		25	45	26	4	0
		20		28	48	22	3	0
		25		31	48	19	2	0
8	3	10		41	51	9	0	0
		15		49	47	4	0	
		20		52	45	3	0	
		25		55	42	2		
		4	10	57	42	2		
		15		62	37	1		
		20		67	32	1		
		25		70	29	0		
		5	10	67	33	0		
		15		74	26	0		
		20		77	23	0		
		25		82	18	0		
7	6	10		74	26	0		
		15		81	19			
		20		84	16			
		25		87	13			
		7	10	80	20			
		15		85	15			
		20		89	11			
		25		91	9			
		8	10	84	16			
		15		90	10			
		20		92	8			
		25		94	6			

**Table 7 Comparison of Tests for 6 Dose Experiments**

POWER COMPARISON OF TESTS FOR 6 DOSE EXPERIMENTS  
LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

REPS	SAMP SIZE	SQRTRULE	TEST	NOAEL					
				0	1	2	3	4	5
2	10	NO	DUNN			15	59	27	
			DUNNETT	6	22	35	27	9	2
			JONCKH	39	47	13	2	0	
		YES	JONCKHX	39	43	17	1		
			MW				100		
			WILLIAMS	59	33	33	13	2	0
	15	NO	DUNN					100	
			DUNNETT	2	14	32	29	18	6
			JONCKH	47	33	16	4	0	
		YES	JONCKHX	46	30	19	4	0	
			MW				100		
			WILLIAMS	57	33	35	15	3	0
20	10	NO	DUNN			16	64	21	
			DUNNETT	8	29	35	22	6	1
			JONCKH	46	45	10	1		
		YES	JONCKHX	48	40	12	1		
			MW				100		
			WILLIAMS	61	40	30	8	1	0
	25	NO	DUNN				100		
			DUNNETT	3	19	36	28	12	3
			JONCKH	53	33	12	3	0	
		YES	JONCKHX	52	31	14	3	0	
			MW				100		
			WILLIAMS	59	39	32	11	1	0
3	10	NO	DUNN			17	66	16	
			DUNNETT	10	30	37	20	4	1
			JONCKH	50	44	6	0		
		YES	JONCKHX	50	41	9	0		
			WILLIAMS	100					
			DUNN				100		
	10	NO	DUNNETT	5	24	36	26	8	2
			JONCKH	57	31	11	2	0	
			JONCKHX	58	28	13	2		
		YES	MW				100		
			WILLIAMS	60	41	30	8	1	0
			DUNN			17	70	14	

**Table 7 Comparison of Tests for 6 Dose Experiments**

POWER COMPARISON OF TESTS FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

REPS	SAMP SIZE	SQR RULE	TEST	NOAEL					
				0	1	2	3	4	5
15	NO	DUNN			5	72	23	1	
		DUNNETT		17	47	32	5	0	
		JONCKH		30	53	17	1		
		JONCKHX		13	59	27	2		
		MW					100		
		WILLIAMS		67	49	16	1		
	YES	DUNN			6	67	27	2	
		DUNNETT		14	50	33	4	0	
		JONCKH		36	50	14	1	0	
		JONCKHX		38	43	19	1		
		MW					100		
		WILLIAMS		68	51	13	1		
20	NO	DUNN			4	76	20	0	
		DUNNETT		18	51	28	3	0	
		JONCKH		31	53	15	1		
		MW					100		
		WILLIAMS		37	50	13	0		
		DUNN			5	69	25	1	
	YES	DUNNETT		16	54	28	3		
		JONCKH		37	51	11	0		
		MW					100		
		WILLIAMS		39	50	10	0		
		DUNN			5	78	18	0	
		DUNNETT		22	52	25	2		
25	NO	JONCKH		37	51	12	0		
		JONCKHX		18	61	20	1		
		MW					100		
		MWX					100		
		WILLIAMS		41	49	11	0		
		DUNN			6	73	21	1	
	YES	DUNNETT		19	55	26	2	0	
		JONCKH		38	52	10	0		
		JONCKHX		38	47	15	0		
		MW					100		
		MWX				2	18	80	
		WILLIAMS		42	51	8	0		
4	10	DUNN			0	29	65	6	0
		DUNNETT		19	53	27	3	0	
		JONCKH		37	48	16	1		
		JONCKHX		23	62	14	1		
		MW					100		
		WILLIAMS		70	48	13	0		
	YES	DUNN			1	51	46	3	0
		DUNNETT		19	55	24	2	0	
		JONCKH		41	50	9	0		
		JONCKHX		38	51	11	0		
		MW		34	37	21	5	2	0
		WILLIAMS		71	50	9	0		
15	NO	DUNN			0	33	65	3	
		DUNNETT		25	57	18	1		
		JONCKH		45	45	11	0		
		JONCKHX		29	60	11			
		MW					100		
		WILLIAMS		73	48	7	0		
	YES	DUNN			0	56	44	1	
		DUNNETT		24	60	16	0		
		JONCKH		47	49	6	0		
		JONCKHX		41	51	8			
		MW		41	40	16	3	0	0
		WILLIAMS		74	48	5	0		

**Table 7 Comparison of Tests for 6 Dose Experiments**

POWER COMPARISON OF TESTS FOR 6 DOSE EXPERIMENTS

LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

REPS	SAMPLESIZE	SQRTRULE	TEST	NOAEL					
				0	1	2	3	4	5
20	NO	DUNN				35	63	2	0
			DUNNETT	28	57	14	0		
			JONCKH	48	45	8	0		
		MW						100	
			WILLIAMS	50	45	5	0		
			DUNN		0	60	38	1	
	YES	DUNNETT		27	60	13	0		
			JONCKH	50	46	4			
			MW	46	39	13	1	0	
		WILLIAMS		52	45	3			
			DUNN			37	62	2	
			DUNNETT	30	58	12	0		
25	NO	JONCKH		49	45	7	0		
			JONCKHX	33	61	7			
		MW					100		
			MW				100		
		WILLIAMS		50	46	4	0		
			DUNN		0	64	36	0	
	YES	DUNNETT		32	60	9	0		
			JONCKH	54	44	3	0		
			JONCKHX	48	48	4			
		MW		51	39	10	1	0	
			MW	47	44	9	1		
			WILLIAMS	56	43	2			
5	10	NO	DUNN		1	61	38	1	
			DUNNETT	26	60	15	0		
			JONCKH	43	50	8	0		
		JONCKHX		42	50	8			
			MW	39	35	19	6	1	0
			WILLIAMS	74	48	5	0		
	YES	DUNN		7	76	18	0		
			DUNNETT	29	59	13	0		
			JONCKH	51	45	5			
		JONCKHX		42	52	6			
			MW	42	44	14	1	0	
			WILLIAMS	76	45	3			
15	NO	DUNN		0	67	33	0		
			DUNNETT	32	59	10	0		
			JONCKH	48	47	6	0		
		JONCKHX		47	47	6	0		
			MW	45	36	16	3	0	
			WILLIAMS	77	44	3	0		
	YES	DUNN		6	82	12	0		
			DUNNETT	34	60	7	0		
			JONCKH	57	41	3			
		JONCKHX		52	45	3			
			MW	47	44	9	0	0	
			WILLIAMS	79	41	2	0		
20	NO	DUNN		0	71	29			
			DUNNETT	36	57	7			
			JONCKH	52	45	3			
		MW		50	36	13	2	0	
			WILLIAMS	57	41	2			
			DUNN		6	85	10	0	
	YES	DUNNETT		37	58	4	0		
			JONCKH	60	38	1			
			MW	52	43	5	0		
		WILLIAMS		62	37	1			

**Table 7 Comparison of Tests for 6 Dose Experiments**

POWER COMPARISON OF TESTS FOR 6 DOSE EXPERIMENTS

LINEAR DOSE-RESPONSE TREND·RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

REPS	SAMP SIZE	SQR RULE	TEST	NOAEL				
				0	1	2	3	4
25	NO	DUNN	DUNN	0	75	25	0	
			DUNNETT	40	56	5	0	
			JONCKH	55	42	3		
			JONCKHX	53	44	3		
			MW	54	35	10	1	0
			MWX				100	
	YES	WILLIAMS	WILLIAMS	61	39	2		
			DUNN		5	88	7	
			DUNNETT	42	55	3	0	
			JONCKH	64	35	1		
			JONCKHX	58	41	1		
			MW	56	40	4	0	
6	10	NO	MWX	57	39	4	0	
			WILLIAMS	66	34	1		
			DUNN		3	80	17	0
			DUNNETT	32	58	10	0	
			JONCKH	48	48	5	0	
			JONCKHX	46	50	4		
	YES	WILLIAMS	MW	47	41	12	1	0
			DUNN	77	44	3	0	
			DUNNETT	0	22	72	7	0
			JONCKH	32	61	8	0	
			JONCKHX	56	42	3		
			MW	54	44	2		
15	NO	WILLIAMS	MW	52	40	8	0	
			WILLIAMS	78	42	2		
			DUNN		2	87	12	0
			DUNNETT	39	57	5	0	
			JONCKH	55	44	2		
			JONCKHX	56	43	2		
	YES	WILLIAMS	MW	54	40	7	0	0
			DUNN	79	40	1		
			DUNNETT		25	72	4	
			JONCKH	38	58	5		
			JONCKHX	62	38	1		
			MW	59	36	4	0	
20	NO	WILLIAMS	WILLIAMS	81	38	1		
			DUNN		2	89	8	
			DUNNETT	43	54	3		
			JONCKH	59	39	1		
			MW	59	36	4	0	
			WILLIAMS	64	36	1		
	YES	WILLIAMS	DUNN		27	71	2	
			DUNNETT	42	55	2		
			JONCKH	65	35	1		
			MW	64	34	2	0	
			WILLIAMS	68	32	0		
			DUNN		2	91	7	
25	NO	WILLIAMS	DUNNETT	45	53	2		
			JONCKH	62	38	1		
			JONCKHX	62	37	1		
			MW	61	36	3	0	100
			MWX				100	
			WILLIAMS	67	34	0		
	YES	WILLIAMS	DUNN		28	71	1	
			DUNNETT	47	51	2		
			JONCKH	69	31	0		
			JONCKHX	66	33	0		
			MW	68	31	2		
			MWX	65	33	2		
			WILLIAMS	72	28	0		

**Table 7 Comparison of Tests for 6 Dose Experiments**

POWER COMPARISON OF TESTS FOR 6 DOSE EXPERIMENTS

LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

REPS	SAMPLESIZE	SQRTRULE	TEST	NOAEL					
				0	1	2	3	4	5
7	NO	DUNN		8	85	8	0		
		DUNNETT		38	57	6	0		
		JONCKH		56	42	3			
		JONCKHX		53	44	3			
		MW		56	38	6	0		
	YES	WILLIAMS		80	39	2			
		DUNN		0	40	59	2		
		DUNNETT		38	59	4			
		JONCKH		61	39	1			
		JONCKHX		56	44	1			
15	NO	MW		60	36	4			
		WILLIAMS		81	37	1			
		DUNN		8	89	4			
		DUNNETT		45	53	2	0		
		JONCKH		63	37	1			
	YES	JONCKHX		61	38	1			
		MW		62	34	4	0		
		WILLIAMS		84	33	0			
		DUNN		0	46	54	1		
		DUNNETT		47	52	2			
20	NO	JONCKH		69	31	0			
		JONCKHX		67	33	0			
		MW		69	30	1			
		WILLIAMS		85	30	0			
		DUNN		7	90	2			
	YES	DUNNETT		51	48	1			
		JONCKH		68	32	0			
		MW		67	30	2			
		WILLIAMS		72	28	0			
		DUNN		0	51	49	0		
25	NO	DUNNETT		53	46	1			
		JONCKH		69	30	0			
		MW		69	30	2	0		
		WILLIAMS		75	25	0			
		DUNN		7	92	1			
	YES	DUNNETT		53	46	1			
		JONCKH		69	30	0			
		MW		76	24	0			
		WILLIAMS		76	24	1			
		DUNN		0	53	47	0		
8	NO	DUNNETT		55	44	1			
		JONCKH		76	24	0			
		MW		76	24	1			
		WILLIAMS		77	23	0			
		DUNN		15	83	3			
	YES	DUNNETT		42	56	3			
		JONCKH		63	36	1			
		JONCKHX		59	40	1			
		MW		58	39	4	0		
		WILLIAMS		83	35	1			

**Table 7 Comparison of Tests for 6 Dose Experiments**

POWER COMPARISON OF TESTS FOR 6 DOSE EXPERIMENTS  
LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

REPS	SAMP SIZE	SQR RULE	TEST	NOAEL				
				0	1	2	3	4
15	NO	DUNN	DUNN	16	83	1		
			DUNNETT	53	47	1		
			JONCKH	70	30	0		
			JONCKHX	67	33	0		
			MW	65	33	2		
	YES	WILLIAMS	WILLIAMS	86	28	0		
			DUNN	0	63	38	0	
			DUNNETT	54	46	1		
			JONCKH	75	26	0		
			JONCKHX	73	27			
20	NO	MW	MW	73	27	0		
			WILLIAMS	88	24	0		
			DUNN	15	84	1		
			DUNNETT	58	41	1		
			JONCKH	75	24	0		
	YES	WILLIAMS	MW	71	28	1		
			DUNN	77	23	0		
			DUNNETT	0	66	33	0	
			JONCKH	60	40	0		
			MW	79	21	0		
25	NO	WILLIAMS	MW	77	23	0		
			WILLIAMS	80	20	0		
			DUNN	80	15	84	0	
			DUNNETT	61	38	0		
			JONCKH	78	22	0		
	YES	MW	MW	74	25	1		
			WILLIAMS	80	20	0		
			DUNN	80	0	71	29	
			DUNNETT	63	63	37	0	
			JONCKH	82	82	18	0	

**Table 8 Comparison of Allocation Rules for 6 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST DUNN	REPS 2	SAMP SIZE	SQR RULE	NOAEL				
				0	1	2	3	4
3	10	NO	NO			15	59	27
			YES				100	
		15	NO			16	64	21
			YES				100	
			NO			17	66	16
	25	YES	NO				100	
			YES			17	70	14
		20	NO				100	
			YES				100	
			NO			17	70	14
4	15	NO	NO			5	64	30
			YES			5	60	33
		20	NO			5	72	23
			YES			6	67	27
			NO			4	76	20
	25	YES	NO			5	69	25
			YES			5	78	18
		20	NO			6	73	21
			YES			0	29	65
			NO			1	51	46
5	20	NO	NO			0	33	65
			YES			0	56	44
		25	NO			35	63	2
			YES			0	60	38
			NO			0	37	62
	25	YES	NO			0	64	36
			YES			1	61	38
		30	NO			7	76	18
			YES			0	67	33
			NO			6	82	12
6	30	NO	NO			0	71	29
			YES			6	85	10
		25	NO			0	75	25
			YES			5	88	7
			NO			3	80	17
	35	YES	NO			0	22	72
			YES			2	87	12
		20	NO			25	72	4
			YES			2	89	8
			NO			27	71	2
7	40	NO	NO			2	91	7
			YES			28	71	1
		35	NO			8	85	8
			YES			0	40	59
			NO			8	89	4
	45	YES	NO			0	46	54
			YES			0	51	49
		30	NO			7	90	2
			YES			0	53	47
			NO			7	92	1
8	50	NO	NO			0	53	47
			YES			0	28	64
		40	NO			16	83	1
			YES			0	63	38
			NO			15	84	1
	45	YES	NO			0	66	33
			YES			15	84	0
		35	NO			0	71	29
			YES					

**Table 8 Comparison of Allocation Rules for 6 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST	REPS	SAMP SIZE	SQRTRULE	NOAEL					
				0	1	2	3	4	5
DUNNETT	2	10	NO	6	22	35	27	9	2
			YES	2	14	32	29	18	6
		15	NO	8	29	35	22	6	1
			YES	3	19	36	28	12	3
		20	NO	10	30	37	20	4	1
			YES	5	24	36	26	8	2
		25	NO	12	33	37	17	3	0
			YES	6	26	38	23	7	1
	3	10	NO	13	42	36	9	1	0
			YES	10	43	38	10	1	
		15	NO	17	47	32	5	0	
			YES	14	50	33	4	0	
		20	NO	18	51	28	3	0	
			YES	16	54	28	3	0	
		25	NO	22	52	25	2	0	
			YES	19	55	26	2	0	
4	4	10	NO	19	53	27	3	0	
			YES	19	55	24	2	0	
		15	NO	25	57	18	1		
			YES	24	60	16	0		
		20	NO	28	57	14	0		
			YES	27	60	13	0		
		25	NO	30	58	12	0		
			YES	32	60	9	0		
		30	NO	26	60	15	0		
			YES	29	59	13	0		
	5	10	NO	32	59	10	0		
			YES	34	60	7	0		
		15	NO	36	57	7			
			YES	37	58	4	0		
		20	NO	40	56	5	0		
			YES	42	55	3	0		
6	6	10	NO	32	58	10	0		
			YES	32	61	8	0		
		15	NO	39	57	5	0		
			YES	38	58	5			
		20	NO	43	54	3			
			YES	42	55	2			
		25	NO	45	53	2			
			YES	47	51	2			
	7	10	NO	38	57	6	0		
			YES	38	59	4			
		15	NO	45	53	2	0		
			YES	47	52	2			
		20	NO	51	48	1			
			YES	53	46	1			
8	8	10	NO	53	46	1			
			YES	55	44	1			
		15	NO	42	56	3			
			YES	38	56	7	0		
	9	15	NO	53	47	1			
			YES	54	46	1			
		20	NO	58	41	1			
			YES	60	40	0			
	25	NO	61	38	0				
			YES	63	37	0			

**Table 8 Comparison of Allocation Rules for 6 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST	REPS	SAMP SIZE	SQRTRULE	NOAEL					
				0	1	2	3	4	5
JONCKH	2	10	NO		39	47	13	2	0
			YES		47	33	16	4	0
		15	NO		46	45	10	1	
			YES		53	33	12	3	0
		20	NO		50	44	6	0	
			YES		57	31	11	2	0
	3	25	NO		52	43	6	0	
			YES		60	30	10	2	0
		10	NO		26	50	23	2	0
			YES		30	50	19	2	0
		15	NO		30	53	17	1	
			YES		36	50	14	1	0
4	4	20	NO		31	53	15	1	
			YES		37	51	11	0	
		25	NO		37	51	12	0	
			YES		38	52	10	0	
		10	NO		37	48	16	1	
			YES		41	50	9	0	
	5	15	NO		45	45	11	0	
			YES		47	49	6	0	
		20	NO		48	45	8	0	
			YES		50	46	4		
		25	NO		49	45	7	0	
			YES		54	44	3	0	
5	5	10	NO		43	50	8	0	
			YES		51	45	5		
		15	NO		48	47	6	0	
			YES		57	41	3		
		20	NO		52	45	3		
			YES		60	38	1		
	6	25	NO		55	42	3		
			YES		64	35	1		
		10	NO		48	48	5	0	
			YES		56	42	3		
		15	NO		55	44	2		
			YES		62	38	1		
6	6	20	NO		59	39	1		
			YES		65	35	1		
		25	NO		62	38	1		
			YES		69	31	0		
		7	NO		56	42	3		
			YES		61	39	1		
	7	15	NO		63	37	1		
			YES		69	31	0		
		20	NO		68	32	0		
			YES		73	26	0		
		25	NO		69	30	0		
			YES		76	24	0		
8	8	10	NO		63	36	1		
			YES		59	39	2		
		15	NO		70	30	0		
			YES		75	26	0		
		20	NO		75	24	0		
			YES		79	21	0		
	8	25	NO		78	22	0		
			YES		82	18	0		

**Table 8 Comparison of Allocation Rules for 6 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST	REPS	SAMP SIZE	SQR RULE	NOAEL				
				0	1	2	3	4
JONCKHX	2	10	NO		39	43	17	1
			YES		46	30	19	4
		15	NO		48	40	12	1
			YES		52	31	14	3
		20	NO		50	41	9	0
			YES		58	28	13	2
		25	NO		51	41	8	0
			YES		60	28	11	1
	3	10	NO		12	53	33	3
			YES		30	43	26	2
		15	NO		13	59	27	2
			YES		38	43	19	1
		25	NO		18	61	20	1
			YES		38	47	15	0
4	4	10	NO		23	62	14	1
			YES		38	51	11	0
		15	NO		29	60	11	
			YES		41	51	8	
		25	NO		33	61	7	
			YES		48	48	4	
		5	NO		42	50	8	
			YES		42	52	6	
		15	NO		47	47	6	0
			YES		52	45	3	
		25	NO		53	44	3	
			YES		58	41	1	
6	6	10	NO		46	50	4	
			YES		54	44	2	
		15	NO		56	43	2	
			YES		61	38	1	
		25	NO		62	37	1	
			YES		66	33	0	
		7	NO		53	44	3	
			YES		56	44	1	
7	7	10	NO		61	38	1	
			YES		67	33	0	
		15	NO		59	40	1	
			YES		47	49	5	
		25	NO		67	33	0	
			YES		73	27		
		10	NO				100	
			YES				100	
MW	2	15	NO				100	
			YES				100	
		20	NO				100	
			YES				100	
		25	NO				100	
			YES				100	
		3	NO				100	
			YES				100	
		15	NO				100	
			YES				100	
		20	NO				100	
			YES				100	
8	4	25	NO				100	
			YES				100	
		10	NO				100	
			YES				100	
		15	NO				100	
			YES				100	
		20	NO				100	
			YES				100	
		25	NO				100	
			YES				100	
		10	NO				100	
			YES		34	37	21	5
		15	NO		41	40	16	3
			YES		46	39	13	1
		20	NO				100	
			YES				100	
		25	NO				100	
			YES		51	39	10	1

**Table 8 Comparison of Allocation Rules for 6 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST	REPS	SAMP SIZE	SQRTRULE	NOAEL					
				0	1	2	3	4	5
5	10	NO		39	35	19	6	1	0
			YES	42	44	14	1	0	
		15	NO	45	36	16	3	0	
			YES	47	44	9	0	0	
		20	NO	50	36	13	2	0	
			YES	52	43	5	0		
			NO	54	35	10	1	0	100
			YES	56	40	4	0		
			NO	47	41	12	1	0	
	6	15	YES	52	40	8	0		
			NO	54	40	7	0	0	
			YES	59	36	4	0		
			NO	59	36	4	0		
			YES	64	34	2	0		
		25	NO	61	36	3	0		100
			YES	68	31	2			
			NO	56	38	6	0		
			YES	60	36	4			
			NO	62	34	4	0		
	7	20	YES	69	30	1			
			NO	67	30	2			
			YES	73	26	1			
			NO	73	26	1			
			YES	69	30	2	0		
		25	NO	76	24	1			
			YES	58	39	4	0		
			NO	67	32	1	0		
			YES	65	33	2			
			NO	73	27	0			
	8	20	NO	71	28	1			
			YES	77	23	0			
			NO	74	25	1			
			YES	80	20	0			
		25	NO					100	
			YES					100	
			NO					100	
			YES				2	18	80
			NO						100
MWX	2	25	YES						
			NO						
			YES						
			NO						
	3	25	NO						
			YES						
			NO						
			YES						
WILLIAMS	2	25	NO						
			YES						
			NO						
			YES						
	3	25	NO						
			YES						
			NO						
			YES						
3	10	25	NO						
			YES						
			NO						
			YES						
	15	25	NO						
			YES						
			NO						
			YES						
20	10	25	NO						
			YES						
			NO						
			YES						
	15	25	NO						
			YES						
			NO						
			YES						
25	10	25	NO						
			YES						
			NO						
			YES						
	15	25	NO						
			YES						
			NO						
			YES						

**Table 8 Comparison of Allocation Rules for 6 Dose Experiments**

POWER COMPARISON OF ALLOCATION RULES FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST	REPS	SAMP SIZE	SQRTRULE	NOAEL				
				0	1	2	3	4
WILLIAMS	3	25	NO	41	49	11	0	
			YES	42	51	8	0	
	4	10	NO	70	48	13	0	
			YES	71	50	9	0	
	5	15	NO	73	48	7	0	
			YES	74	48	5	0	
	6	20	NO	50	45	5	0	
			YES	52	45	3		
	7	25	NO	50	46	4	0	
			YES	56	43	2		
4	10	NO	74	48	5	0		
			YES	76	45	3		
	15	NO	77	44	3	0		
			YES	79	41	2	0	
	20	NO	57	41	2			
			YES	62	37	1		
	25	NO	61	39	2			
			YES	66	34	1		
5	10	NO	77	44	3	0		
			YES	78	42	2		
	15	NO	79	40	1			
			YES	81	38	1		
	20	NO	64	36	1			
			YES	68	32	0		
	25	NO	67	34	0			
			YES	72	28	0		
6	10	NO	80	39	2			
			YES	81	37	1		
	15	NO	84	33	0			
			YES	85	30	0		
	20	NO	72	28	0			
			YES	75	25	0		
	25	NO	73	27	0			
			YES	77	23	0		
7	10	NO	83	35	1			
			YES	85	29	0		
	15	NO	86	28	0			
			YES	88	24	0		
	20	NO	77	23	0			
			YES	80	20	0		
	25	NO	80	20	0			
			YES	83	17	0		

**Table 9 Effect of Number of Replicates on Power**

EFFECT OF NUMBER OF REPLICATES ON POWER FOR 6 DOSE EXPERIMENTS  
 LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)  
 MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST DUNN	SQRTRULE NO	REPS 2	SAMPSIZE	NOAEL				
				0	1	2	3	4
			10			15	59	27
			15			16	64	21
			20			17	66	16
			25			17	70	14
		3	10		5	64	30	2
		3	15		5	72	23	1
		3	20		4	76	20	0
		3	25		5	78	18	0
		4	10	0	29	65	6	0
		4	15	0	33	65	3	
		4	20		35	63	2	0
		4	25		37	62	2	
		5	10	1	61	38	1	
		5	15	0	67	33	0	
		5	20	0	71	29		
		5	25	0	75	25	0	
		6	10	3	80	17	0	
		6	15	2	87	12	0	
		6	20	2	89	8		
		6	25	2	91	7		
		7	10	8	85	8	0	
		7	15	8	89	4		
		7	20	7	90	2		
		7	25	7	92	1		
		8	10	15	83	3		
		8	15	16	83	1		
		8	20	15	84	1		
		8	25	15	84	0		
YES	2		10				100	
YES	2		15				100	
YES	2		20				100	
YES	2		25				100	
	3		10		5	60	33	3
	3		15		6	67	27	2
	3		20		5	69	25	1
	3		25		6	73	21	1
	4		10	1	51	46	3	0
	4		15	0	56	44	1	
	4		20	0	60	38	1	
	4		25	0	64	36	0	
	5		10	7	76	18	0	
	5		15	6	82	12	0	
	5		20	6	85	10	0	
	5		25	5	88	7		
	6		10	0	22	72	7	0
	6		15		25	72	4	
	6		20		27	71	2	
	6		25		28	71	1	
	7		10	0	40	59	2	
	7		15	0	46	54	1	
	7		20	0	51	49	0	
	7		25	0	53	47	0	
	8		10	0	28	64	9	0
	8		15	0	63	38	0	
	8		20	0	66	33	0	
	8		25	0	71	29		

**Table 9 Effect of Number of Replicates on Power**

EFFECT OF NUMBER OF REPLICATES ON POWER FOR 6 DOSE EXPERIMENTS

LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST	SQRTRULE	REPS	SAMPSIZE	NOAEL					
				0	1	2	3	4	5
DUNNETT	NO	2	10	6	22	35	27	9	2
			15	8	29	35	22	6	1
			20	10	30	37	20	4	1
			25	12	33	37	17	3	0
			3	10	13	42	36	9	1
	3	10	15	17	47	32	5	0	
			20	18	51	28	3	0	
			25	22	52	25	2		
			4	10	19	53	27	3	0
			15	25	57	18	1		
YES	2	10	20	28	57	14	0		
			25	30	58	12	0		
			5	10	26	60	15	0	
			15	32	59	10	0		
			20	36	57	7			
	3	10	25	40	56	5	0		
			15	32	58	10	0		
			20	39	57	5	0		
			25	43	54	3			
			6	10	45	53	2		
8	2	10	10	38	57	6	0		
			15	45	53	2	0		
			20	51	48	1			
			25	53	46	1			
			8	10	42	56	3		
	3	10	15	53	47	1			
			20	58	41	1			
			25	61	38	0			
			10	2	14	32	29	18	6
			15	3	19	36	28	12	3
YES	4	10	20	5	24	36	26	8	2
			25	6	26	38	23	7	1
			3	10	10	43	38	10	1
			15	14	50	33	4	0	
			20	16	54	28	3		
	5	10	25	19	55	26	2	0	
			15	24	60	16	0		
			20	27	60	13	0		
			25	32	60	9	0		
			6	10	29	59	13	0	
8	6	10	15	34	60	7	0		
			20	37	58	4	0		
			25	42	55	3	0		
			6	10	32	61	8	0	
			15	38	58	5			
	7	10	20	42	55	2			
			25	47	51	2			
			7	10	38	59	4		
			15	47	52	2			
			20	53	46	1			
8	10	25	25	55	44	1			
			10	38	56	7	0		
			15	54	46	1			
			20	60	40	0			
			25	63	37	0			

**Table 9 Effect of Number of Replicates on Power**

EFFECT OF NUMBER OF REPLICATES ON POWER FOR 6 DOSE EXPERIMENTS

LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST JONCKH	SQRTRULE NO	REPS 2	SAMP SIZE	NOAEL					
				0	1	2	3	4	5
3	NO	10		39	47	13	2	0	
		15		46	45	10	1		
		20		50	44	6	0		
		25		52	43	6	0		
		10		26	50	23	2	0	
	YES	15		30	53	17	1		
		20		31	53	15	1		
		25		37	51	12	0		
		10		37	48	16	1		
		15		45	45	11	0		
4	NO	20		48	45	8	0		
		25		49	45	7	0		
		10		43	50	8	0		
		15		48	47	6	0		
		20		52	45	3			
	YES	25		55	42	3			
		10		48	48	5	0		
		15		55	44	2			
		20		59	39	1			
		25		62	38	1			
5	NO	10		56	42	3			
		15		63	37	1			
		20		68	32	0			
		25		69	30	0			
		10		63	36	1			
	YES	15		70	30	0			
		20		75	24	0			
		25		78	22	0			
		10		47	33	16	4	0	
		15		53	33	12	3	0	
6	NO	20		57	31	11	2	0	
		25		60	30	10	2	0	
		10		30	50	19	2	0	
		15		36	50	14	1	0	
		20		37	51	11	0		
	YES	25		38	52	10	0		
		10		41	50	9	0		
		15		47	49	6	0		
		20		50	46	4			
		25		54	44	3	0		
7	NO	10		51	45	5			
		15		57	41	3			
		20		60	38	1			
		25		64	35	1			
		10		56	42	3			
	YES	15		62	38	1			
		20		65	35	1			
		25		69	31	0			
		10		61	39	1			
		15		69	31	0			
8	NO	20		73	26	0			
		25		76	24	0			
		10		59	39	2			
		15		75	26	0			
		20		79	21	0			
	YES	25		82	18	0			

**Table 9 Effect of Number of Replicates on Power**

EFFECT OF NUMBER OF REPLICATES ON POWER FOR 6 DOSE EXPERIMENTS

LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST JONCKHX	SQRTRULE NO	REPS 2	SAMP SIZE	NOAEL					
				0	1	2	3	4	5
3	NO	10		39	43	17	1		
		15		48	40	12	1		
		20		50	41	9	0		
		25		51	41	8	0		
	YES	10		12	53	33	3		
		15		13	59	27	2		
		25		18	61	20	1		
	NO	10		23	62	14	1		
		15		29	60	11			
		25		33	61	7			
	YES	10		42	50	8			
		15		47	47	6	0		
		25		53	44	3			
4	NO	10		46	50	4			
		15		56	43	2			
		25		62	37	1			
		10		53	44	3			
	YES	15		61	38	1			
		10		59	40	1			
		15		67	33	0			
	NO	10		46	30	19	4	0	
		15		52	31	14	3	0	
		20		58	28	13	2		
	YES	25		60	28	11	1		
		10		30	43	26	2		
		15		38	43	19	1		
5	NO	25		38	47	15	0		
		10		38	51	11	0		
		15		41	51	8			
		25		48	48	4			
	YES	10		42	52	6			
		15		52	45	3			
		25		58	41	1			
	NO	10		54	44	2			
		15		61	38	1			
		25		66	33	0			
	YES	10		56	44	1			
		15		67	33	0			
		10		47	49	5			
	NO	15		73	27				
6	NO	2					100		
		10					100		
		15					100		
		25					100		
	YES	10					100		
		15					100		
		20					100		
	NO	25					100		
		10					100		
		15					100		
	YES	20					100		
		25					100		
		10		39	35	19	6	1	0
7	NO	15		45	36	16	3	0	
		20		50	36	13	2	0	
		25		54	35	10	1	0	100
		10		47	41	12	1	0	
	YES	15		54	40	7	0	0	
		20		59	36	4	0		
		25		61	36	3	0		100
	NO	10		56	38	6	0		
		15		62	34	4	0		
		20		67	30	2			
	YES	25		69	30	2	0		

*Appendix 1 to the proposal for Phase 2 of the validation of the amphibian metamorphosis assay*

8	10	58	39	4	0			
	15	65	33	2				
	20	71	28	1				
	25	74	25	1				
YES	2	10				100		
	15					100		
	20					100		
	25					100		
	3	10				100		
	15					100		
	20					100		
	25					100		
	4	10	34	37	21	5	2	0
		15	41	40	16	3	0	0
		20	46	39	13	1	0	
		25	51	39	10	1	0	
	5	10	42	44	14	1	0	
		15	47	44	9	0	0	
		20	52	43	5	0		
		25	56	40	4	0		
	6	10	52	40	8	0		
		15	59	36	4	0		
		20	64	34	2	0		
		25	68	31	2			
	7	10	60	36	4			
		15	69	30	1			
		20	73	26	1			
		25	76	24	1			
	8	10	67	32	1	0		
		15	73	27	0			
		20	77	23	0			
		25	80	20	0			
MWX	NO	2	25				100	
		3	25				100	
		4	25				100	
		5	25				100	
		6	25				100	
YES	2	25					100	
		3	25			2	18	80
		4	25	47	44	9	1	
		5	25	57	39	4	0	
		6	25	65	33	2		
WILLIAMS NO	2	10	59	33	33	13	2	0
		15	61	40	30	8	1	0
		20				100		
		25	26	44	26	6	0	
	3	10	65	48	21	2	0	
		15	67	49	16	1		
		20	37	50	13	0		
		25	41	49	11	0		
	4	10	70	48	13	0		
		15	73	48	7	0		
		20	50	45	5	0		
		25	50	46	4	0		
	5	10	74	48	5	0		
		15	77	44	3	0		
WILLIAMS NO	5	20	57	41	2			
		25	61	39	2			
	6	10	77	44	3	0		
		15	79	40	1			
		20	64	36	1			
		25	67	34	0			
	7	10	80	39	2			
		15	84	33	0			
		20	72	28	0			
		25	73	27	0			
	8	10	83	35	1			
		15	86	28	0			
		20	77	23	0			
		25	80	20	0			

**Table 9 Effect of Number of Replicates on Power**

EFFECT OF NUMBER OF REPLICATES ON POWER FOR 6 DOSE EXPERIMENTS

LINEAR DOSE-RESPONSE TREND RELATIVE TO LOG(DOSE)

MAX EFFECT=0.8754, EFFECTS=0 0.17508 0.35016 0.52524 0.70032 0.8754

TEST	SQRTRULE	REPS	SAMP SIZE	NOAEL					
				0	1	2	3	4	5
YES	2	10	57	33	35	15	3	0	
		15	59	39	32	11	1	0	
		20	60	41	30	8	1	0	
		25	22	42	30	7	0		
		10	65	50	19	1	0		
	3	15	68	51	13	1			
		20	39	50	10	0			
		25	42	51	8	0			
		10	71	50	9	0			
4	5	15	74	48	5	0			
		20	52	45	3				
		25	56	43	2				
		10	76	45	3				
		15	79	41	2	0			
	6	20	62	37	1				
		25	66	34	1				
		10	78	42	2				
		15	81	38	1				
		20	68	32	0				
8	7	25	72	28	0				
		10	81	37	1				
		15	85	30	0				
		20	75	25	0				
		25	77	23	0				
	8	10	85	29	0				
		15	88	24	0				
		20	80	20	0				
		25	83	17	0				