# On the Comparison of Umbrella Pattern Treatment Means with a Control Mean

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#### Summary

This paper is concerned with comparing several increasing dose levels (treatments) with a zero dose control when the prior information about the umbrella pattern treatment means is available. The problem of testing whether there is at least one treatment which is better than the control is considered. Multiple test procedures are then proposed for deciding treatments (if any) which are better than the control. Some approximate criticial values of the proposed tests are reported. The results of a Monte Carlo power study are presented.

Key words: Monte Carlo study; Multiple test procedure; Umbrella pattern treatment means.

### 1. Introduction

The problem of comparing increasing dose levels (treatments) of a substance with a zero dose control together with the prior information that if there were a response to the substance the treatment means would be monotonically ordered was first considered in Williams (1971, 1972). However, monotonicity of dose-response relationship is far from universal. Many examples are available in medicine where increasing doses of therapies usually produce better (say, higher) treatment effects, but these therapies often become counter-productive at high doses. In such cases, an increasing dose-response relationship with a downturn in response at high doses is anticipated. These treatment effects are then expected to have an up-down ordering, hence, they are said to follow an umbrella pattern (see, for instance, MACK and WOLFE, 1981). The point which separates the treatment effects into the two different ordering groups is called the peak of the umbrella.

Suppose that  $X_{i1}, \ldots, X_{in_i}$ ,  $i=0,1,\ldots,k$  are k+1 independent random samples from normal distributions  $N(\mu_0, \sigma^2), N(\mu_1, \sigma^2), \ldots, N(\mu_k, \sigma^2)$ , where  $\mu_0$  represents the control mean and  $\mu_i$  the *i*th treatment mean, and  $\sigma^2$  is the common variance. Specifically, we consider testing the null hypothesis  $H_0: [\mu_0 = \mu_1 = \ldots = \mu_k]$  against the alternative hypothesis  $H_a: [\mu_i > \mu_0$  for at least one i]. In addition, we assume that, under  $H_a, \mu_1 \leq \ldots \leq \mu_p \geq \ldots \geq \mu_k$ , for some p, with at least one strict inequality.

In section 2 we propose an extension of Williams' test to the case of known umbrella peak. Estimation of the doses (if any) which are better than the control is also discussed. In section 3 we propose a test procedure for the case where the unknown peak group is expected to be relatively close to the kth population. A testing procedure for the more general unknown peak setting based on the method of estimating the umbrella peak suggested by Shi (1988) is also considered. In section 4 we report some approximate critical values of the tests proposed in sections 2 and 3, respectively. In section 5 we present the results of an extensive Monte Carlo simulation investigation of the relative powers of several competing tests for a variety of umbrella pattern treatment means configurations.

## 2. Case of Known Umbrella Peak

Suppose that, under  $H_A$ , the peak of the umbrella known to be at group p  $(1 \le p \le k)$ . Further, assume that  $n_0 = c$  and  $n_1 = \ldots = n_k = n$ . Let  $\overline{X}_0, \overline{X}_1, \ldots, \overline{X}_k$  be the sample means for the k+1 groups and let  $\hat{\mu}_1^{(p)} \le \ldots \le \hat{\mu}_p^{(p)} \ge \ldots \ge \hat{\mu}_k^{(p)}$  be the isotonic regression of  $\overline{X}_1, \ldots, \overline{X}_k$  under the restrictions  $\mu_1 \le \ldots \le \mu_p \ge \ldots \ge \mu_k$ . Note that if  $\mu_p > \mu_0$ , then  $H_A$  holds. Therefore, we propose to reject  $H_0$  for large values of

$$T_{p} = (\hat{\mu}_{p}^{(p)} - \bar{X}_{0})/(S \sqrt{n^{-1} + c^{-1}}), \tag{2.1}$$

where  $S^2$  is an unbiased estimator of the common variance  $\sigma^2$  independent of  $\bar{X}_0, \bar{X}_1, \ldots, \bar{X}_k$  and  $vS^2/\sigma^2$  is distributed to chi-squared distribution with degrees of freedom v = k(n-1) + c - 1. Note that the test based on  $T_k$  is in fact Williams' test for comparing ordered treatment means with a control mean.

From the derivation of the isotonic regression  $\hat{\mu}_1^{(p)} \leq ... \leq \hat{\mu}_p^{(p)} \geq ... \geq \hat{\mu}_k^{(p)}$  discussed in Chen and Wolfe (1990), we obtain

$$\hat{\mu}_p^{(p)} = \max_{1 \le s \le p \le t \le k} \sum_{i=s}^t \bar{X}_i / (t - s + 1). \tag{2.2}$$

Therefore, the statistic  $T_p$  becomes

$$T_{p} = \max_{1 \le s \le p \le t \le k} \left\{ \sum_{i=s}^{t} \overline{X}_{i} / (t - s + 1) - \overline{X}_{0} \right\} / (S \sqrt{n^{-1} + c^{-1}}). \tag{2.3}$$

Hence, under  $H_0$ , the distribution of  $T_p$  is the same as that of  $(Y-Z_0)/W^{1/2}$ , where  $Y = \max_{1 \le s \le p \le t \le k} \sum_{i=s}^t Z_i/(t-s+1)$ , W and the  $Z_i$  are independent random variables,  $Z_0$  is distributed according to N(0, n/c),  $Z_1, \ldots, Z_k$  are standard normal random variables and cvW is distributed as  $(c+n)\chi_v^2$ .

If the test based on  $T_p$  rejects the null hypothesis  $H_0$ , we conclude that there is a response to the substance and that at least the dose p is better than the control. However, it is possible that this response occurs not only at dose p. Let  $t(\alpha; p, k)$  be the upper  $\alpha$ th percentile for the null distribution of  $T_p$ . Set  $T_{pi} = (\hat{\mu}_i^{(p)} - \bar{X}_0)/S \sqrt{n^{-1} + c^{-1}}, i = 1, ..., k$ . If  $T_{pa} < t(\alpha; k, p), T_{pb} < t(\alpha; k, p)$  and  $T_{pi} \ge t(\alpha; k, p)$  for  $1 \le a and <math>a < i < b$ , we then conclude that there is evidence for a response at doses a + 1, ..., b - 1. Since

$$\alpha = P\{T_p \ge t(\alpha; k, p) | H_0\} \ge P\{T_{pi} \ge t(\alpha; k, p), i = 1, ..., k | H_0\},$$

in the language of multiple comparison procedures, the experiment-wise error rate of the procedure stated above is controlled.

## 3. Case of Unknown Umbrella Peak

When treatment effects follow an umbrella pattern, it occurs frequently that this umbrella has a downturn at high dose levels (see, for example, SIMPSON and MARGOLIN (1986)). In this case, the peak group of the umbrella is expected to be relatively close to the kth population. Hence, we suggest to choose the peak group  $\tilde{p}$  such that

$$\tilde{p} = \max \left\{ i \colon \bar{X}_i > \frac{\bar{X}_1 + \dots + \bar{X}_{i-1}}{i-1} \right\}.$$

Finally, the null hypothese  $H_0$  is rejected for large values of

$$T_{\tilde{p}} = \max_{1 \le s \le \tilde{p} \le t \le k} \left\{ \sum_{i=s}^{t} \bar{X}_i / (t-s+1) - \bar{X}_0 \right\} / (S \sqrt{n^{-1} + c^{-1}}). \tag{3.1}$$

For the more general setting where the peak of the umbrella is unknown, we employ the method suggested by SHI (1988) to estimate the unknown peak. Let  $\hat{\mu}_1^{(t)} \leq \ldots \leq \hat{\mu}_t^{(t)} \geq \ldots \geq \hat{\mu}_k^{(t)}$  be the isotonic regression of  $\overline{X}_1, \ldots, \overline{X}_k$  under the restrictions  $\mu_1 \leq \ldots \leq \mu_t \geq \ldots \geq \mu_k$  for  $t = 1, \ldots, k$ . We first select the group  $\hat{p}$  such that

$$\sum_{i=1}^{k} (\hat{\mu}_{i}^{(\hat{p})} - \bar{X}_{i})^{2} = \min_{1 \le t \le k} \sum_{i=1}^{k} (\hat{\mu}_{i}^{(t)} - \bar{X}_{i})^{2}.$$

The null hypothesis  $H_0$  is then rejected for large values of

$$T_{\hat{p}} = \max_{1 \le s \le \hat{p} \le t \le k} \left\{ \sum_{i=s}^{t} \bar{X}_i / (t - s + 1) - \bar{X}_0 \right\} / (S \sqrt{n^{-1} + c^{-1}}). \tag{3.2}$$

Let  $t_{\tilde{p}}(\alpha; k)$  and  $t_{\hat{p}}(\alpha; k)$  be the upper  $\alpha$ th percentiles for the null distributions of  $T_{\tilde{p}}$  and  $T_{\hat{p}}$ , respectively. If the test based on  $T_{\tilde{p}}$  (or  $T_{\hat{p}}$ ) rejects  $H_0$ , a level

 $(1-\alpha)$  multiple comparison procedure similar to that described in section 2, but employing the critical value  $t_{\tilde{p}}(\alpha; k)$  (or  $t_{\hat{p}}(\alpha; k)$ ), can be used to estimate doses which are significantly better than the zero-dose control.

# 4. Null Distributions of the Proposed Tests

When, in particular, k = 3, p = 2, n = c, and the common variance  $\sigma^2$  is known, we observe from section 2 that the statistic  $T_2$  is distributed as

$$\max_{1 \le s \le 2 \le t \le 3} \left\{ \sum_{i=s}^{t} Z_i / (t-s+1) - Z_0 \right\} / \sqrt{2},$$

where the  $Z_i$  are independent standard normal random variables. According to the 6 different permutations of  $Z_1$ ,  $Z_2$  and  $Z_3$ , we obtain

$$\begin{split} \sqrt{2}\,T_2 &= \bar{Z}_{23} - Z_0, & \text{if } Z_2 < Z_3 \text{ and } \bar{Z}_{23} > Z_1 \\ &= \bar{Z}_{12} - Z_0, & \text{if } Z_1 > Z_2 \text{ and } \bar{Z}_{12} > Z_3 \\ &= Z_2 - Z_0, & \text{if } Z_1 < Z_2 \text{ and } Z_2 > Z_3 \\ &= \bar{Z}_{123} - Z_0, & \text{if } Z_1 < Z_3 \text{ and } \bar{Z}_{23} < Z_1 \text{ or } Z_1 > Z_3 \text{ and } \bar{Z}_{12} < Z_3, \end{split}$$

where  $\bar{Z}_{st} = (Z_s + Z_t)/2$  for  $1 \le s \le 2 \le t \le 3$  and  $\bar{Z}_{123} = (Z_1 + Z_2 + Z_3)/3$ . After some algebraic manipulations, we have

$$\begin{split} P\{T_2 &\geq t \,|\, H_0\} = \int\limits_0^\infty \exp\left(-x^2/12\right) \,\varPhi\left\{(x-6\,t)/4\,\sqrt{3}\right\} dx/2\,\sqrt{3\,\pi} \\ &+ \int\limits_0^\infty \int\limits_0^\infty \exp\left\{-(2\,y-x)^2/12 - x^2/4\right\} \,\varPhi\left\{(3\,t-x-y)/2\,\sqrt{3}\right\} dx\,dy/2\,\sqrt{3\,\pi} \\ &+ \varPhi\left(\sqrt{3}\,t/2\right) \int\limits_0^\infty \exp\left(-x^2/4\right) \,\varPhi\left(-3\,x/\sqrt{6}\right) dx/\sqrt{\pi} \,. \end{split}$$

By using elementary numerical integration techniques to solve the equation  $P\{T_2 \ge t(\alpha; 3, 2) | H_0\} = \alpha$ , we obtain the critical values  $t(\alpha; 3, 2)$  for  $\alpha = .05$  and .10 in the following:

Table 1 Critical values  $t(\alpha; 3, 2)$  of the test based on  $T_2$  when n = c and  $\sigma^2$  is known

| α                 | .05    | .10    |
|-------------------|--------|--------|
| $t(\alpha; 2, 3)$ | 1.7697 | 1.4402 |

Since the null distribution of the test statistic  $T_p$  becomes relatively complicated for  $k \ge 4$  and that of the statistic  $T_{\tilde{p}}$  is even more difficult to obtain, we employ simulation technique to estimate the critical values  $t(\alpha; k, p)$  and  $t_{\tilde{p}}(\alpha; k)$ . For each value of k (and p for peak-known case), the number of treatments, the International Mathematical and Statistical Libraries (IMSL) routine RNNOR was used to generate appropriate normal random variates for which the statistics  $T_p$  and  $T_{\tilde{p}}$  were respectively evaluated. Proceeding in this fashion, we obtained empirical cumulative distributions of  $T_p$  and  $T_{\tilde{p}}$ , respectively, based on a sample of size 10,000 from the corresponding true distributions. The estimated critical values for the  $T_p$  and  $T_{\tilde{p}}$  tests then correspond to percentiles of their empirical distributions and are presented in Tables 2 and 3. When k=3 with infinite degrees of freedom, for example, the estimated 95th percentile for the null distribution of  $T_{\tilde{p}}$  is 1.917.

Table 2 Estimated critical values of  $T_n$  for  $n_0 = n_1 = ... = n_k$ 

|          |   |       |       |       |       | (a) ( | $\alpha = 0.05$ |       |       |       |       |
|----------|---|-------|-------|-------|-------|-------|-----------------|-------|-------|-------|-------|
|          | k |       | 3     | 80    | 4     | 8     | 5               |       |       | 6     |       |
| d. f.    | p | 2     | 3     | 3     | 4     | 3     | 4               | 5     | 4     | 5     | 6     |
| 5        |   | 2.285 | 2.247 | 2.252 | 2.228 | 2.346 | 2.346           | 2.242 | 2.292 | 2.262 | 2.192 |
| 6        |   | 2.163 | 2.083 | 2.172 | 2.101 | 2.200 | 2.187           | 2.192 | 2.210 | 2.185 | 2.145 |
| 7        |   | 2.165 | 2.111 | 2.153 | 2.104 | 2.129 | 2.112           | 2.029 | 2.139 | 2.101 | 2.074 |
| 8        |   | 2.033 | 1.991 | 2.067 | 2.050 | 2.092 | 2.076           | 2.026 | 2.069 | 2.089 | 2.024 |
| 9        |   | 2.066 | 2.012 | 2.043 | 1.990 | 2.048 | 2.061           | 1.990 | 2.064 | 2.054 | 1.992 |
| 10       |   | 1.989 | 1.981 | 1.976 | 1.938 | 1.985 | 2.024           | 1.945 | 2.050 | 2.041 | 1.967 |
| 11       |   | 2.051 | 1.946 | 1.977 | 1.942 | 2.008 | 2.053           | 1.976 | 2.035 | 2.041 | 1.930 |
| 12       |   | 1.932 | 1.894 | 1.980 | 1.884 | 2.005 | 1.982           | 1.922 | 1.983 | 1.966 | 1.913 |
| 13       |   | 1.943 | 1.888 | 1.956 | 1.902 | 1.998 | 2.008           | 1.942 | 1.972 | 1.994 | 1.912 |
| 14       |   | 1.945 | 1.861 | 1.950 | 1.892 | 1.948 | 1.923           | 1.893 | 1.958 | 1.933 | 1.912 |
| 15       |   | 1.931 | 1.893 | 1.938 | 1.890 | 1.981 | 1.993           | 1.947 | 1.967 | 1.965 | 1.919 |
| 16       |   | 1.913 | 1.847 | 1.879 | 1.877 | 1.905 | 1.909           | 1.851 | 1.978 | 1.957 | 1.914 |
| 17       |   | 1.910 | 1.877 | 1.913 | 1.864 | 1.936 | 1.957           | 1.905 | 1.937 | 1.949 | 1.895 |
| 18       |   | 1.890 | 1.866 | 1.897 | 1.864 | 1.929 | 1.948           | 1.891 | 1.944 | 1.945 | 1.889 |
| 19       |   | 1.879 | 1.845 | 1.887 | 1.857 | 1.913 | 1.938           | 1.874 | 1.940 | 1.932 | 1.884 |
| 20       |   | 1.860 | 1.832 | 1.878 | 1.849 | 1.911 | 1.931           | 1.873 | 1.941 | 1.928 | 1.894 |
| 22       |   | 1.852 | 1.813 | 1.867 | 1.837 | 1.907 | 1.909           | 1.849 | 1.931 | 1.919 | 1.876 |
| 24       |   | 1.848 | 1.813 | 1.860 | 1.824 | 1.904 | 1.916           | 1.844 | 1.912 | 1.913 | 1.875 |
| 26       |   | 1.852 | 1.815 | 1.855 | 1.823 | 1.903 | 1.910           | 1.845 | 1.896 | 1.892 | 1.853 |
| 28       |   | 1.848 | 1.798 | 1.861 | 1.827 | 1.895 | 1.898           | 1.828 | 1.897 | 1.882 | 1.842 |
| 30       |   | 1.839 | 1.787 | 1.858 | 1.815 | 1.881 | 1.878           | 1.813 | 1.898 | 1.891 | 1.857 |
| 35       |   | 1.836 | 1.796 | 1.846 | 1.804 | 1.883 | 1.873           | 1.805 | 1.874 | 1.870 | 1.828 |
| 40       |   | 1.828 | 1.770 | 1.834 | 1.792 | 1.894 | 1.876           | 1.815 | 1.876 | 1.867 | 1.826 |
| 60       |   | 1.816 | 1.756 | 1.821 | 1.775 | 1.850 | 1.832           | 1.787 | 1.859 | 1.847 | 1.800 |
| 120      |   | 1.795 | 1.744 | 1.825 | 1.751 | 1.833 | 1.817           | 1.775 | 1.823 | 1.841 | 1.797 |
| $\infty$ |   | 1.770 | 1.730 | 1.779 | 1.740 | 1.771 | 1.778           | 1.740 | 1.795 | 1.802 | 1.760 |

Table 2 (continued)

|          |   |       |       |       |       | (b)   | $\alpha = 0.10$ |       |       |       |       |
|----------|---|-------|-------|-------|-------|-------|-----------------|-------|-------|-------|-------|
| 2 2      | k |       | 3     |       | 4     |       | 5               |       |       | 6     |       |
| v. f.    | p | 2     | 3     | 3     | 4     | 3     | 4               | 5     | 4     | 5     | 6     |
| 5        |   | 1.729 | 1.684 | 1.767 | 1.696 | 1.775 | 1.786           | 1.699 | 1.779 | 1.756 | 1.679 |
| 6        |   | 1.638 | 1.635 | 1.679 | 1.614 | 1.727 | 1.715           | 1.663 | 1.731 | 1.694 | 1.635 |
| 7        |   | 1.657 | 1.609 | 1.672 | 1.626 | 1.672 | 1.658           | 1.595 | 1.686 | 1.657 | 1.629 |
| 8        |   | 1.596 | 1.548 | 1.624 | 1.583 | 1.646 | 1.641           | 1.560 | 1.655 | 1.656 | 1.613 |
| 9        |   | 1.620 | 1.557 | 1.601 | 1.567 | 1.639 | 1.619           | 1.565 | 1.632 | 1.621 | 1.564 |
| 10       |   | 1.560 | 1.521 | 1.576 | 1.516 | 1.604 | 1.615           | 1.574 | 1.626 | 1.616 | 1.553 |
| 11       |   | 1.576 | 1.535 | 1.560 | 1.529 | 1.616 | 1.614           | 1.569 | 1.617 | 1.601 | 1.531 |
| 12       |   | 1.532 | 1.472 | 1.578 | 1.534 | 1.590 | 1.595           | 1.533 | 1.566 | 1.557 | 1.514 |
| 13       |   | 1.543 | 1.506 | 1.547 | 1.507 | 1.594 | 1.585           | 1.550 | 1.603 | 1.586 | 1.529 |
| 14       |   | 1.547 | 1.495 | 1.559 | 1.507 | 1.542 | 1.529           | 1.485 | 1.563 | 1.560 | 1.508 |
| 15       |   | 1.544 | 1.509 | 1.537 | 1.505 | 1.587 | 1.594           | 1.552 | 1.584 | 1.578 | 1.507 |
| 16       |   | 1.508 | 1.476 | 1.520 | 1.472 | 1.544 | 1.539           | 1.492 | 1.592 | 1.567 | 1.523 |
| 17       |   | 1.519 | 1.495 | 1.524 | 1.500 | 1.574 | 1.573           | 1.529 | 1.569 | 1.566 | 1.516 |
| 18       |   | 1.510 | 1.487 | 1.511 | 1.486 | 1.572 | 1.559           | 1.516 | 1.573 | 1.564 | 1.515 |
| 19       |   | 1.496 | 1.476 | 1.504 | 1.486 | 1.569 | 1.556           | 1.506 | 1.572 | 1.561 | 1.507 |
| 20       |   | 1.493 | 1.471 | 1.497 | 1.484 | 1.561 | 1.544           | 1.504 | 1.565 | 1.563 | 1.503 |
| 22       |   | 1.488 | 1.461 | 1.490 | 1.469 | 1.550 | 1.531           | 1.491 | 1.565 | 1.557 | 1.493 |
| 24       |   | 1.483 | 1.453 | 1.478 | 1.462 | 1.546 | 1.541           | 1.487 | 1.553 | 1.542 | 1.492 |
| 26       |   | 1.482 | 1.451 | 1.477 | 1.451 | 1.538 | 1.541           | 1.484 | 1.542 | 1.527 | 1.475 |
| 28       |   | 1.471 | 1.442 | 1.482 | 1.459 | 1.529 | 1.535           | 1.478 | 1.541 | 1.531 | 1.477 |
| 30       |   | 1.478 | 1.445 | 1.476 | 1.450 | 1.521 | 1.527           | 1.475 | 1.534 | 1.537 | 1.484 |
| 35       |   | 1.480 | 1.448 | 1.466 | 1.444 | 1.524 | 1.518           | 1.466 | 1.527 | 1.522 | 1.476 |
| 40       |   | 1.469 | 1.436 | 1.459 | 1.441 | 1.532 | 1.532           | 1.470 | 1.525 | 1.521 | 1.482 |
| 60       |   | 1.460 | 1.430 | 1.446 | 1.436 | 1.495 | 1.453           | 1.553 | 1.502 | 1.503 | 1.470 |
| 120      |   | 1.449 | 1.420 | 1.460 | 1.427 | 1.479 | 1.493           | 1.447 | 1.489 | 1.505 | 1.465 |
| $\infty$ |   | 1.440 | 1.390 | 1.444 | 1.419 | 1.460 | 1.461           | 1.419 | 1.481 | 1.475 | 1.434 |

Table 3 Estimated critical values of  $T_{\tilde{p}}$  for  $n_0 = n_1 = ... = n_k$ 

| d.f. |   |       | (a) α | =0.05 |       |       | (b) α | (b) $\alpha = 0.10$ |       |  |
|------|---|-------|-------|-------|-------|-------|-------|---------------------|-------|--|
| v    | k | 3     | 4     | 5     | 6     | 3     | 4     | 5                   | 6     |  |
| 5    |   | 2.515 | 2.633 | 2.719 | 2.711 | 1.953 | 2.075 | 2.122               | 2.118 |  |
| 6    |   | 2.342 | 2.509 | 2.613 | 2.606 | 1.825 | 1.976 | 2.045               | 2.057 |  |
| 7    |   | 2.387 | 2.450 | 2.481 | 2.490 | 1.863 | 1.975 | 1.965               | 2.014 |  |
| 8    |   | 2.237 | 2.373 | 2.401 | 2.442 | 1.786 | 1.913 | 1.960               | 1.991 |  |
| 9    |   | 2.230 | 2.330 | 2.377 | 2.414 | 1.786 | 1.884 | 1.916               | 1.945 |  |
| 10   |   | 2.200 | 2.294 | 2.330 | 2.351 | 1.765 | 1.841 | 1.910               | 1.911 |  |
| 11   |   | 2.175 | 2.249 | 2.353 | 2.349 | 1.744 | 1.837 | 1.905               | 1.926 |  |
| 12   |   | 2.116 | 2.242 | 2.291 | 2.283 | 1.710 | 1.817 | 1.852               | 1.852 |  |
| 13   |   | 2.115 | 2.206 | 2.284 | 2.301 | 1.716 | 1.824 | 1.876               | 1.908 |  |
| 14   |   | 2.120 | 2.201 | 2.227 | 2.257 | 1.720 | 1.805 | 1.809               | 1.865 |  |

Table 3 (continued)

| d.f.     |   |       | (a) α | =0.05 |       |       | (b) α | = 0.10 |       |
|----------|---|-------|-------|-------|-------|-------|-------|--------|-------|
| v        | k | 3     | 4     | 5     | 6     | 3     | 4     | 5      | 6     |
| 15       |   | 2.107 | 2.192 | 2.260 | 2.271 | 1.712 | 1.800 | 1.871  | 1.888 |
| 16       |   | 2.070 | 2.146 | 2.214 | 2.248 | 1.682 | 1.748 | 1.813  | 1.869 |
| 17       |   | 2.079 | 2.174 | 2.218 | 2.254 | 1.698 | 1.789 | 1.838  | 1.870 |
| 18       |   | 2.065 | 2.159 | 2.202 | 2.245 | 1.681 | 1.780 | 1.828  | 1.870 |
| 19       |   | 2.039 | 2.149 | 2.188 | 2.244 | 1.666 | 1.771 | 1.816  | 1.857 |
| 20       |   | 2.028 | 2.134 | 2.190 | 2.245 | 1.655 | 1.762 | 1.811  | 1.863 |
| 22       |   | 2.015 | 2.113 | 2.173 | 2.229 | 1.646 | 1.752 | 1.792  | 1.850 |
| 24       |   | 2.012 | 2.098 | 2.167 | 2.220 | 1.642 | 1.745 | 1.796  | 1.840 |
| 26       |   | 2.018 | 2.101 | 2.168 | 2.197 | 1.640 | 1.742 | 1.794  | 1.826 |
| 28       |   | 1.999 | 2.110 | 2.156 | 2.183 | 1.632 | 1.745 | 1.785  | 1.818 |
| 30       |   | 1.999 | 2.103 | 2.131 | 2.197 | 1.629 | 1.736 | 1.780  | 1.824 |
| 35       |   | 1.985 | 2.080 | 2.123 | 2.164 | 1.633 | 1.724 | 1.775  | 1.808 |
| 40       |   | 1.961 | 2.071 | 2.127 | 2.167 | 1.624 | 1.711 | 1.780  | 1.803 |
| 60       |   | 1.946 | 2.054 | 2.079 | 2.135 | 1.616 | 1.692 | 1.761  | 1.786 |
| 120      |   | 1.913 | 2.036 | 2.065 | 2.112 | 1.595 | 1.683 | 1.748  | 1.775 |
| $\infty$ |   | 1.917 | 2.045 | 2.038 | 2.073 | 1.605 | 1.700 | 1.708  | 1.745 |

# 5. Monte Carlo Power Study

To examine the relative powers of the proposed tests based on  $T_p$ ,  $T_{\tilde{p}}$  and  $T_{\hat{p}}$ , Shi's tests based on  $\bar{\chi}_p^2$  and  $\bar{\chi}_{\hat{p}}^2$ , Williams' test based on W and Dunnett's test based on D for comparing umbrella pattern treatment means with a control mean, we conducted a Monte Carlo Study. In this study, we consider k=4 treatments, with known  $\sigma^2$  and c/n=1 and 3, where  $n_0=c$  and  $n_1=\ldots=n_k=n$ , and a variety of different umbrella pattern treatment means.

The power performances of the test procedures considered in this paper are evaluated via the experiment-wise power (probability of detecting at least one treatment which is better than the control) and the comparison-wise power (probability of correctly detecting the treatments which are better than the control, available only for multiple test procedures). For each of these settings, appropriate normal variates were generated by using the IMSL routines RNNOR. In each case, we used 10,000 replications in obtaining the various power estimates. Approximate level  $\alpha = 0.05$  and 0.10 critical values were used. The simulated power estimates for the seven tests are presented in Tables 4 and 5. The designated umbrella pattern treatment means correspond to values of  $\mu_{10} = \mu_1 - \mu_0, \dots, \mu_{k0} = \mu_k - \mu_0$ .

We observe from the simulation results that Williams' test has excellent experiment-wise power when the treatment means have a monotonic ordering. Likewise, the test based on  $T_p$  provides excellent experiment-wise power against umbrella pattern treatment means when the peak is correctly chosen. This is not

Table 4 Experiment-wise power estimates for k = 4 when  $\sigma^2$  is known

|            |            |            |            | (a) | $\alpha = 0$ . | 05   |                  |                     |               |                          |      |
|------------|------------|------------|------------|-----|----------------|------|------------------|---------------------|---------------|--------------------------|------|
| $\mu_{10}$ | $\mu_{20}$ | $\mu_{30}$ | $\mu_{40}$ | c/n | $T_p$          | W    | $\bar{\chi}_p^2$ | $T_{\widetilde{p}}$ | $T_{\hat{p}}$ | $\bar{\chi}^2_{\hat{p}}$ | D    |
| 0          | 0          | 0          | 2.4        | 1   | .493           | .493 | .518             | .371                | .326          | .336                     | .325 |
|            |            |            |            | 3   | .637           | .637 | .568             | .515                | .463          | .372                     | .463 |
| 0          | 0          | 1.2        | 2.4        | 1   | .509           | .509 | .567             | .390                | .350          | .384                     | .350 |
|            |            |            |            | 3   | .661           | .661 | .641             | .544                | .502          | .454                     | .502 |
| 0          | 1.2        | 1.2        | 2.4        | 1   | .513           | .513 | .522             | .401                | .371          | .356                     | .371 |
|            |            |            |            | 3   | .669           | .669 | .654             | .562                | .536          | .476                     | .536 |
| 1.2        | 2.4        | 2.4        | 3.6        | 1   | .826           | .826 | .786             | .743                | .726          | .643                     | .725 |
|            |            |            |            | 3   | .951           | .951 | .949             | .918                | .912          | .884                     | .912 |
| 0          | 0          | 2.4        | 0          | 1   | .472           | .153 | .512             | .337                | .322          | .393                     | .322 |
|            |            |            |            | 3   | .633           | .198 | .548             | .466                | .457          | .428                     | .457 |
| 0          | 1.2        | 2.4        | 0          | 1   | .491           | .170 | .551             | .373                | .347          | .432                     | .346 |
|            |            |            |            | 3   | .660           | .228 | .618             | .525                | .498          | .508                     | .498 |
| 0          | 1.2        | 2.4        | 1.2        | 1   | .505           | .326 | .502             | .370                | .369          | .387                     | .368 |
|            |            |            |            | 3   | .682           | .464 | .613             | .525                | .533          | .507                     | .533 |
| 1.2        | 2.4        | 3.6        | 1.2        | 1   | .812           | .525 | .769             | .721                | .695          | .666                     | .694 |
|            |            |            |            | 3   | .948           | .747 | .915             | .893                | .884          | .863                     | .884 |
| 0          | 2.4        | 0          | 0          | 1   | .480           | .090 | .511             | .311                | .320          | .401                     | .319 |
|            |            |            |            | 3   | .649           | .098 | .547             | .422                | .456          | .442                     | .456 |
| 0          | 2.4        | 1.2        | 0          | 1   | .496           | .140 | .542             | .344                | .346          | .430                     | .345 |
|            |            |            |            | 3   | .674           | .186 | .609             | .475                | .496          | .508                     | .496 |
| 1.2        | 2.4        | 1.2        | 0          | 1   | .510           | .153 | .516             | .389                | .367          | .413                     | .366 |
|            |            |            |            | 3   | .694           | .207 | .633             | .545                | .531          | .535                     | .531 |
| 1.2        | 3.6        | 2.4        | 1.2        | 1   | .808           | .491 | .760             | .688                | .699          | .670                     | .698 |
|            |            |            |            | 3   | .948           | .716 | .904             | .870                | .884          | .859                     | .884 |
| 2.4        | 0          | 0          | 0          | 1   | .482           | .072 | .512             | .101                | .323          | .387                     | .322 |
|            |            |            |            | 3   | .647           | .074 | .530             | .123                | .456          | .432                     | .456 |
| 2.4        | 1.2        | 0          | 0          | 1   | .500           | .098 | .550             | .213                | .348          | .426                     | .347 |
|            |            |            |            | 3   | .671           | .106 | .593             | .299                | .496          | .503                     | .496 |
| 2.4        | 1.2        | 1.2        | 0          | 1   | .505           | .144 | .500             | .262                | .369          | .390                     | .367 |
|            |            |            |            | 3   | .679           | .184 | .590             | .377                | .528          | .517                     | .528 |
| 3.6        | 2.4        | 2.4        | 1.2        | 1   | .821           | .501 | .739             | .635                | .722          | .646                     | .721 |
|            |            |            |            | 3   | .956           | .733 | .914             | .851                | .909          | .883                     | .909 |

| (b) $\alpha = 0.10$ |            |            |            |     |       |      |                  |                     |               |                          |      |
|---------------------|------------|------------|------------|-----|-------|------|------------------|---------------------|---------------|--------------------------|------|
| $\mu_{10}$          | $\mu_{20}$ | $\mu_{30}$ | $\mu_{40}$ | c/n | $T_p$ | W    | $\bar{\chi}_p^2$ | $T_{\widetilde{p}}$ | $T_{\hat{p}}$ | $\bar{\chi}^2_{\hat{p}}$ | D    |
| 0                   | 0          | 0          | 2.4        | 1   | .614  | .614 | .652             | .510                | .458          | .464                     | .456 |
|                     |            |            |            | 3   | .755  | .755 | .695             | .637                | .584          | .499                     | .584 |
| 0                   | 0          | 1.2        | 2.4        | 1   | .631  | .631 | .694             | .533                | .490          | .516                     | .488 |
|                     |            |            |            | 3   | .785  | .785 | .760             | .670                | .631          | .583                     | .630 |
| 0                   | 1.2        | 1.2        | 2.4        | 1   | .639  | .639 | .660             | .549                | .516          | .488                     | .515 |
|                     |            |            |            | 3   | .795  | .795 | .774             | .693                | .669          | .609                     | .668 |

Table 4 (continued)

|            |            |            |            | (b) | $\alpha = 0.$ | 10   |                  |                     |                   |                          |      |
|------------|------------|------------|------------|-----|---------------|------|------------------|---------------------|-------------------|--------------------------|------|
| $\mu_{10}$ | $\mu_{20}$ | $\mu_{30}$ | $\mu_{40}$ | c/n | $T_p$         | W    | $\bar{\chi}_p^2$ | $T_{\widetilde{p}}$ | $T_{\widehat{p}}$ | $\bar{\chi}^2_{\hat{p}}$ | D    |
| 1.2        | 2.4        | 2.4        | 3.6        | 1   | .899          | .899 | .872             | .850                | .834              | .760                     | .833 |
|            |            |            |            | 3   | .982          | .982 | .978             | .962                | .959              | .936                     | .959 |
| 0          | 0          | 2.4        | 0          | 1   | .610          | .262 | .645             | .468                | .451              | .524                     | .450 |
|            |            |            |            | 3   | .754          | .332 | .687             | .591                | .582              | .561                     | .582 |
| 0          | 1.2        | 2.4        | 0          | 1   | .631          | .294 | .681             | .517                | .486              | .569                     | .485 |
|            |            |            |            | 3   | .781          | .379 | .749             | .660                | .632              | .636                     | .631 |
| 0          | 1.2        | 2.4        | 1.2        | 1   | .647          | .470 | .627             | .516                | .512              | .522                     | .511 |
|            |            |            |            | 3   | .803          | .629 | .743             | .670                | .671              | .634                     | .670 |
| 1.2        | 2.4        | 3.6        | 1.2        | 1   | .894          | .680 | .859             | .833                | .814              | .784                     | .813 |
|            |            |            |            | 3   | .978          | .872 | .960             | .948                | .941              | .922                     | .941 |
| 0          | 2.4        | 0          | 0          | 1   | .602          | .161 | .644             | .436                | .456              | .535                     | .455 |
|            |            |            |            | 3   | .754          | .186 | .681             | .539                | .582              | .573                     | .581 |
| 0          | 2.4        | 1.2        | 0          | 1   | .622          | .246 | .675             | .486                | .491              | .572                     | .489 |
|            |            |            |            | 3   | .782          | .317 | .733             | .615                | .631              | .639                     | .630 |
| 1.2        | 2.4        | 1.2        | 0          | 1   | .638          | .268 | .657             | .538                | .516              | .549                     | .515 |
|            |            |            |            | 3   | .802          | .351 | .757             | .685                | .669              | .666                     | .668 |
| 1.2        | 3.6        | 2.4        | 1.2        | 1   | .888          | .635 | .851             | .806                | .812              | .783                     | .811 |
|            |            |            |            | 3   | .975          | .840 | .950             | .934                | .941              | .917                     | .941 |
| 2.4        | 0          | 0          | 0          | 1   | .620          | .143 | .641             | .195                | .455              | .521                     | .454 |
|            |            |            |            | 3   | .759          | .151 | .659             | .228                | .582              | .563                     | .582 |
| 2.4        | 1.2        | 0          | 0          | 1   | .640          | .188 | .675             | .342                | .486              | .564                     | .485 |
|            |            |            |            | 3   | .789          | .221 | .718             | .441                | .626              | .630                     | .626 |
| 2.4        | 1.2        | 1.2        | 0          | 1   | .647          | .265 | .631             | .407                | .512              | .528                     | .511 |
|            |            |            |            | 3   | .798          | .334 | .720             | .536                | .664              | .642                     | .664 |
| 3.6        | 2.4        | 2.4        | 1.2        | 1   | .901          | .665 | .832             | .777                | .833              | .763                     | .832 |
|            |            |            |            | 3   | .982          | .868 | .956             | .928                | .956              | .932                     | .956 |

Table 5 Comparison-wise power estimates for k = 4 when  $\sigma^2$  is known

| (a) $\alpha = 0.05$ |            |            |            |     |       |      |                     |               |      |  |
|---------------------|------------|------------|------------|-----|-------|------|---------------------|---------------|------|--|
| $\mu_{10}$          | $\mu_{20}$ | $\mu_{30}$ | $\mu_{40}$ | c/n | $T_p$ | W    | $T_{\widetilde{p}}$ | $T_{\hat{p}}$ | D    |  |
| 0                   | 0          | 0          | 2.4        | 1   | .444  | .441 | .347                | .301          | .286 |  |
|                     |            |            |            | 3   | .593  | .588 | .494                | .437          | .424 |  |
| )                   | 0          | 1.2        | 2.4        | 1   | .140  | .135 | .083                | .053          | .049 |  |
|                     |            |            |            | 3   | .194  | .194 | .114                | .068          | .066 |  |
| )                   | 1.2        | 1.2        | 2.4        | 1   | .087  | .086 | .045                | .028          | .018 |  |
|                     |            |            |            | 3   | .111  | .116 | .052                | .030          | .019 |  |
| 1.2                 | 2.4        | 2.4        | 3.6        | 1   | .160  | .177 | .085                | .062          | .044 |  |
|                     |            |            |            | 3   | .199  | .221 | .107                | .069          | .046 |  |
| )                   | 0          | 2.4        | 0          | 1   | .404  | .000 | .276                | .289          | .282 |  |
|                     |            |            |            | 3   | .560  | .000 | .397                | .424          | .419 |  |
| )                   | 1.2        | 2.4        | 0          | 1   | .116  | .000 | .069                | .050          | .050 |  |
|                     |            |            |            | 3   | .173  | .000 | .100                | .066          | .066 |  |

Table 5 (continued)

|            | (a) $\alpha = 0.05$ |            |            |     |       |      |                     |               |      |  |  |
|------------|---------------------|------------|------------|-----|-------|------|---------------------|---------------|------|--|--|
| $\mu_{10}$ | $\mu_{20}$          | $\mu_{30}$ | $\mu_{40}$ | c/n | $T_p$ | W    | $T_{\widetilde{p}}$ | $T_{\hat{p}}$ | D    |  |  |
| 0          | 1.2                 | 2.4        | 1.2        | 1   | .058  | .110 | .037                | .022          | .020 |  |  |
|            |                     |            |            | 3   | .072  | .152 | .041                | .024          | .022 |  |  |
| 1.2        | 2.4                 | 3.6        | 1.2        | 1   | .072  | .176 | .035                | .021          | .018 |  |  |
|            |                     |            |            | 3   | .078  | .222 | .040                | .027          | .024 |  |  |
| 0          | 2.4                 | 0          | 0          | 1   | .412  | .000 | .251                | .288          | .281 |  |  |
|            |                     |            |            | 3   | .573  | .000 | .358                | .425          | .419 |  |  |
| 0          | 2.4                 | 1.2        | 0          | 1   | .120  | .000 | .090                | .049          | .049 |  |  |
|            |                     |            |            | 3   | .185  | .000 | .131                | .066          | .066 |  |  |
| 1.2        | 2.4                 | 1.2        | 0          | 1   | .059  | .000 | .032                | .019          | .017 |  |  |
|            |                     |            |            | 3   | .075  | .000 | .032                | .022          | .020 |  |  |
| 1.2        | 3.6                 | 2.4        | 1.2        | 1   | .073  | .186 | .036                | .021          | .018 |  |  |
|            |                     |            |            | 3   | .083  | .238 | .040                | .027          | .024 |  |  |
| 2.4        | 0                   | 0          | 0          | 1   | .434  | .000 | .000                | .295          | .281 |  |  |
|            |                     |            |            | 3   | .600  | .000 | .000                | .429          | .417 |  |  |
| 2.4        | 1.2                 | 0          | 0          | 1   | .136  | .000 | .147                | .051          | .048 |  |  |
|            |                     |            |            | 3   | .198  | .000 | .217                | .068          | .066 |  |  |
| 2.4        | 1.2                 | 1.2        | 0          | 1   | .089  | .000 | .050                | .026          | .017 |  |  |
|            |                     |            |            | 3   | .120  | .000 | .059                | .027          | .018 |  |  |
| 3.6        | 2.4                 | 2.4        | 1.2        | 1   | .159  | .468 | .096                | .066          | .048 |  |  |
|            |                     |            |            | 3   | .212  | .670 | .116                | .074          | .049 |  |  |

| (b) $\alpha = 0.10$ |            |            |            |     |       |      |                     |               |      |  |  |
|---------------------|------------|------------|------------|-----|-------|------|---------------------|---------------|------|--|--|
| $\mu_{10}$          | $\mu_{20}$ | $\mu_{30}$ | $\mu_{40}$ | c/n | $T_p$ | W    | $T_{\widetilde{p}}$ | $T_{\hat{p}}$ | D    |  |  |
| 0                   | 0          | 0          | 2.4        | 1   | .519  | .513 | .453                | .404          | .376 |  |  |
|                     |            |            |            | 3   | .663  | .656 | .591                | .534          | .506 |  |  |
| )                   | 0          | 1.2        | 2.4        | 1   | .202  | .192 | .143                | .099          | .091 |  |  |
|                     |            |            |            | 3   | .285  | .278 | .187                | .124          | .117 |  |  |
| )                   | 1.2        | 1.2        | 2.4        | 1   | .149  | .138 | .088                | .060          | .042 |  |  |
|                     |            |            |            | 3   | .197  | .192 | .101                | .061          | .039 |  |  |
| 1.2                 | 2.4        | 2.4        | 3.6        | 1   | .258  | .289 | .166                | .123          | .094 |  |  |
|                     |            |            |            | 3   | .322  | .359 | .190                | .139          | .096 |  |  |
| )                   | 0          | 2.4        | 0          | 1   | .477  | .000 | .354                | .383          | .370 |  |  |
|                     |            |            |            | 3   | .613  | .000 | .467                | .518          | .504 |  |  |
| )                   | 1.2        | 2.4        | 0          | 1   | .171  | .000 | .119                | .087          | .087 |  |  |
|                     |            |            |            | 3   | .246  | .000 | .161                | .113          | .113 |  |  |
| )                   | 1.2        | 2.4        | 1.2        | 1   | .105  | .171 | .071                | .046          | .042 |  |  |
|                     |            |            |            | 3   | .129  | .242 | .079                | .048          | .045 |  |  |
| 1.2                 | 2.4        | 3.6        | 1.2        | 1   | .142  | .290 | .085                | .052          | .046 |  |  |
|                     |            |            |            | 3   | .154  | .359 | .079                | .061          | .055 |  |  |
| )                   | 2.4        | 0          | 0          | 1   | .473  | .000 | .316                | .389          | .373 |  |  |
|                     |            |            |            | 3   | .612  | .000 | .412                | .520          | .505 |  |  |

Table 5 (continued)

| $\mu_{10}$ | $\mu_{20}$ | $\mu_{30}$ | $\mu_{40}$ | c/n | $T_p$ | W    | $T_{\widetilde{p}}$ | $T_{\widehat{p}}$ | D    |
|------------|------------|------------|------------|-----|-------|------|---------------------|-------------------|------|
| 0          | 2.4        | 1.2        | 0          | 1   | .170  | .000 | .145                | .090              | .089 |
|            |            |            |            | 3   | .248  | .000 | .200                | .117              | .116 |
| 1.2        | 2.4        | 1.2        | 0          | 1   | .101  | .000 | .065                | .044              | .040 |
|            |            |            |            | 3   | .133  | .000 | .070                | .047              | .043 |
| 1.2        | 3.6        | 2.4        | 1.2        | 1   | .135  | .302 | .086                | .052              | .046 |
|            |            |            |            | 3   | .153  | .378 | .080                | .060              | .054 |
| 2.4        | 0          | 0          | 0          | 1   | .525  | .000 | .000                | .400              | .371 |
|            |            |            |            | 3   | .665  | .000 | .000                | .531              | .503 |
| 2.4        | 1.2        | 0          | 0          | 1   | .204  | .000 | .234                | .099              | .089 |
|            |            |            |            | 3   | .287  | .000 | .320                | .124              | .117 |
| 2.4        | 1.2        | 1.2        | 0          | 1   | .147  | .000 | .100                | .060              | .039 |
|            |            |            |            | 3   | .205  | .000 | .118                | .060              | .041 |
| 3.6        | 2.4        | 2.4        | 1.2        | 1   | .263  | .626 | .172                | .128              | .098 |
|            |            |            |            | 3   | .334  | .826 | .202                | .141              | .099 |

surprising since both tests are designed to detect for their respective special classes of alternatives. However, the power of Williams' test drops sharply when there is a downturn in the umbrella. Similarly, we would expect the power of the test based on  $T_p$  to decline when the peak of the umbrella in incorrectly selected. In addition, the comparison-wise power performance of Williams' procedure is similar to that of the multiple test procedure proposed in section 2 for comparing ordered treatment means with a control mean. For comparing umbrella pattern treatment means with a control mean, however, Williams' procedure may not be as powerful as the proposed procedure, especially for the case where the kth treatment mean of this umbrella is the same as the control mean.

Simulation results indicate that with equal treatment replication Shi's tests based on  $\bar{\chi}_p^2$  and  $\bar{\chi}_{\hat{p}}^2$ , respectively, are better than the corresponding  $T_p$  and  $T_{\hat{p}}$  tests when some of the  $\mu_{i0}$  are zero. When treatments are all better than the control, however, the tests based on  $T_p$  and  $T_{\hat{p}}$  compare more favorably with the corresponding  $\bar{\chi}_p^2$  and  $\bar{\chi}_p^2$  tests. In particular, when the control replication is increased, the tests based on  $T_p$  and  $T_{\hat{p}}$  are superior to  $\bar{\chi}_p^2$  and  $\bar{\chi}_{\hat{p}}^2$  tests, respectively. Moreover, Shi's tests for umbrella alternatives only provide single tests. In comparing several umbrella pattern treatment means with a control mean, however, experimenters usually prefer procedures which can be used to determine which treatments (if any) are more effective than the control.

In general, the statistic  $T_{\tilde{p}}$  provides a better test procedure than does either  $T_{\hat{p}}$  or D for the unknown peak setting when the peak group is relatively close to the kth population. When, however, the location of the peak group is relatively far away from the kth population, the procedure based on  $T_{\tilde{p}}$  performs poorly.

In these cases, the test procedures based on  $T_{\hat{p}}$  and D, respectively, are both superior to the one based on  $T_{\tilde{p}}$ . Finally, we observe that the power performance of the procedure based on  $T_{\hat{p}}$  is similar to that of Dunnett's procedure based on D for comparing general umbrella pattern treatment means with a control mean.

According to the simulation results, we, therefore, have several recommendations. When the prior information that the treatment means have an umbrella pattern under the alternative is available, the procedure based on  $T_p$  should be used if one is relatively confident of the location of the peak group. The procedure based on  $T_{\tilde{p}}$  is recommended if the peak group of the umbrella is unknown, but is believed to be relatively close to the kth population. For the case where no information about the location of the peak group is available, Dunnett's procedure is suggested since it is computationally less complicated than the power-equivalent procedure based on  $T_{\hat{p}}$ .

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