

# Tree-based Multiple Hypothesis Testing with General FWER Control

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March 2, 2025

## Contents

1	Mathematical Formulation	1
2	Weak FWER Control	2
3	General FWER/FDR Control	3
4	Miscellanea	4

## 1. Mathematical Formulation

*Here I consider a testing procedure without data splitting, and only about a case on the 'multiplicative trees'.*

In the tree-structured multiple hypotheses setting, we say  $\mathbf{X}_t$  is a data set to construct each  $t$ th p-value.

Let  $p_t$  be the p-value of the parent node and  $p_{t+1}$  be the p-value of the subsequent child of that parent node. When we fix some  $\alpha \in (0, 1]$ , our basic observation is

$$P(p_{t+1} > \alpha | p_t \leq \alpha_1) \leq P(p_{t+1} > \alpha | p_t \leq \alpha_2),$$

whenever  $\alpha_1 \leq \alpha_2$ . This is because, we can understand our procedure is actually using

$$p'_{t+1} = \max(p_1, \dots, p_t, p_{t+1}),$$

as a  $p$ -value implemented in the  $t+1$ th node, where  $p_i$  is all other ancestors of the  $p_{t+1}$ . This can be another (maybe equivalent) understanding of the **monotonicity condition** in Goeman and Solari (2010).

From this we can recall the **positive regression dependency on each one from a subset  $I_0$  (or PRDS on  $I_0$ )** from Benjamini and Yekutieli (2001), which states when  $D$  is a non-decreasing set (so whenever  $x \in D$  and  $x \leq y$ , then  $y \in D$ ),

$$P(\mathbf{X} \in D | X_i = X)$$

is non-decreasing in  $x$ .

We also provide some notation for the tree-structured hypotheses. Let us have a tree with depth of  $d$  where for some fixed integer  $k$ ,  $k$  is a number of children for every parent node (so we consider *multiplicative trees*). Then we say  $N$  is the number of total nodes and  $n$  as a number of total clusters, where

$$N = \sum_{r=1}^d k^{r-1}$$
$$n = 1 + \sum_{r=1}^{d-1} k^{r-1}.$$

## 2. Weak FWER Control

Here we consider about the weak FWER control, so when we assume all of the nulls are true. We first impose an assumption, called 'conditional super-uniformity', following Robertson et al. (2023). For the  $p$ -values  $p_i$ , we define  $R_i = \mathbb{1}(p_i \leq \alpha_i)$ , where  $\mathbb{F}^t = \sigma(R_1, \dots, R_t)$  and we let  $\alpha_t$  as a  $\mathbb{F}^{t-1}$ -measurable function of  $(R_1, \dots, R_{t-1})$ .

**Definition 1.** The null  $p$ -values are said to be **conditionally super-uniform** if  $P(p_t \leq \alpha_t | \mathbb{F}^{t-1}) \leq \alpha_t$  for any  $\mathbb{F}^{t-1}$ -measurable  $\alpha_t$ .

We also give additional notation before proposing our theorem on the weak FWER control. We say  $C_i$  is an ordered set of parents of the  $i$ th node on the tree, and  $d_i$  is the depth of the  $i$ th node. Then we see  $|C_i| = d_i + 1$ . Denote  $C_{i,j}$  as the  $j$ th element of  $C_i$ . For example, consider a binary tree and the case  $i = 8$ . Then  $C_8 = \{1, 2, 4\}$ , and  $C_{8,2} = 4$ . We are also able to check  $|C_8| = 3$ , and  $d_8 = 2$ .

Now we state our theorem.

**Theorem 1.** If the null  $p$ -values are conditionally super-uniform, then the procedure with the critical value functions for every  $t$ th node

$$\alpha_t = \begin{cases} \frac{\alpha}{1+k\alpha}, & \text{if all of the } i\text{th tests where } i \in C_t \text{ are rejected,} \\ 0, & \text{otherwise,} \end{cases}$$

implies that the weak FWER is controlled at level  $\alpha$ .

*Proof.* Assume that the all of the nulls are true. First, note that for any  $i$ th node and  $\alpha \in (0, 1)$ , since we assumed conditional super-uniformity on the null  $p$ -values,

$$\begin{aligned} P(p_i \leq \alpha) &= P(p_i \leq \alpha \text{ and } \forall j \in C_i, p_j \leq \alpha) + P(p_i \leq \alpha \text{ and } \exists j \in C_i \text{ such that } p_j > \alpha) \\ &= P(p_i \leq \alpha \text{ and } \forall j \in C_i, p_j \leq \alpha) \\ &= P(p_1 \leq \alpha)P(p_{C_{i,2}} \leq \alpha | p_1 \leq \alpha) \dots P(p_i \leq \alpha | p_1 \leq \alpha, \dots, p_{C_{i,d_i}} \leq \alpha) \\ &\leq \alpha^{d_i+1} \end{aligned}$$

holds. Therefore,

$$\begin{aligned} P(V \geq 1) &= P\left(\left(\cap_{i=1}^N \{p_i > \alpha_i\}\right)^c\right) \\ &= P\left(\cup_{i=1}^N \{p_i \leq \alpha_i\}\right) \\ &\leq \sum_{i=1}^N P(p_i \leq \alpha_i) \\ &= \underbrace{P(p_1 \leq \alpha_1)}_{p\text{-value on the root node}} + \underbrace{P(p_2 \leq \alpha_2) + \dots + P(p_{1+k} \leq \alpha_{1+k})}_{k \text{ } p\text{-values on the second level}} + \underbrace{P(p_{2+k} \leq \alpha_{2+k}) + P(p_{1+k+k^2} \leq \alpha_{1+k+k^2})}_{k^2 \text{ } p\text{-values on the third level}} + \dots \\ &\quad + \underbrace{P(p_{N-k^{d-1}+1} \leq \alpha_{N-k^{d-1}+1}) + \dots + P(p_N \leq \alpha_N)}_{k^{d-1} \text{ } p\text{-values on the final level}} \\ &\leq \sum_{j=1}^d k^{j-1} \alpha_j^j \\ &\leq \sum_{j=1}^{\infty} k^{j-1} \left(\frac{\alpha}{1+k\alpha}\right)^j = \alpha, \end{aligned}$$

which gives us a desired conclusion. □

### 3. General FWER/FDR Control

Now we assume that, whenever  $j$  is a child node of  $i$  in the tree structure,  $\mathbf{X}_j \subseteq \mathbf{X}_i$ . Note that this is a much more realistic than the conditional super-uniformity assumption.

## 4. Miscellanea

My idea is that,

- Since  $p_1, \dots, p_n$  (so the  $p$ -values after local adjustments) are still has a PRDS property, we can try to directly apply Benjamini and Yekutieli (2001)'s approach to get general FDR or FWER control.
- We can also consider which kind of local adjustments give valid and more powerful tests.

I will further check about,

- Idea of data splitting used in the recent simulation.
- Prove explicitly the validness of locally adjusted  $p$ -values from different methods.

## REFERENCES

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