

Weighted Adaptive Multiple Decision Functions for False Discovery Rate Control

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Outline

- 1 Introduction: Motivation and FDR Research Areas
- 2 Framework and Exact Results
- 3 Asymptotic FDP control
- 4 Optimal Weights
- 5 Assessment
- 6 Concluding Remarks

Motivation

Introduction: Motivation of Multiple Testing and FDR Research Areas

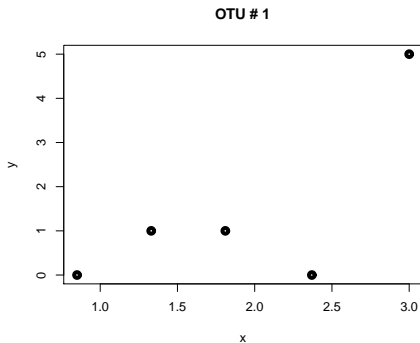
Data: Anderson and Habiger (2012)

Biology Theory: Ecosystem of micro-organisms (OTUs) near the roots of wheat (stomachs of wheat plant).

Question: Which species (OTUs) are associated with productivity?

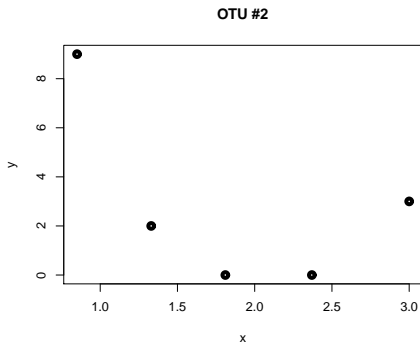
OTU #	Productivity/Shoot Biomass (g)				
	0.85	1.33	1.81	2.37	3.00
1	0	1	1	0	5
2	9	2	0	0	3
⋮	⋮	⋮	⋮	⋮	⋮
M = 778	16	10	29	18	13

Goal 1



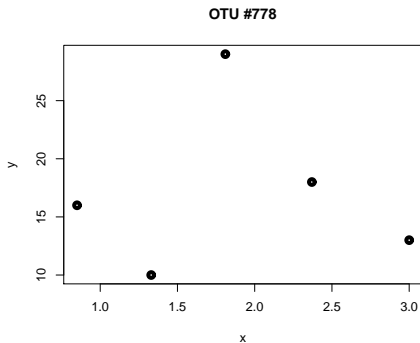
Y (prevalence) vs. X (productivity)?

Goal 2



Y (prevalence) vs. X (productivity)?

Goal 778



Y (prevalence) vs. X (productivity)?

Model

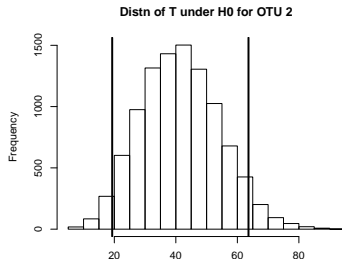
- Data: Y_{mj} prevalence of m th OTU in j th productivity group
- Model:

$$Y_{mj} \sim \text{Poisson}(e^{\beta_{0m} + \beta_{1m}x_j})$$

- Hypotheses: $H_{0m} : \beta_{1m} = 0$ vs. $H_{1m} : \beta_{1m} \neq 0$
- Sufficient Stat: $T_m = \sum_{j=1}^5 x_j Y_{mj}$
- Ancillary Stat: $Y_{m\cdot} = \sum_{j=1}^5 Y_{mj}$

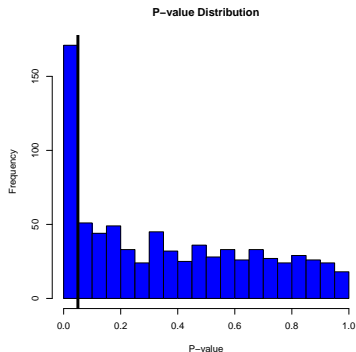
A single test

- $Y_{m1}, Y_{m2}, \dots, Y_{m5} | Y_{m\cdot} \sim \text{Multi}(Y_{m\cdot}, 1/5)$ under H_{m0}



- $T_2 = 19.31 \rightarrow \text{p-value} = 0.050.$ ^T
- Reject $H_{02} : \beta_{12} = 0?$

Multiple Tests



Using p-value cutoff $\alpha = 0.05$...

- **171 Discoveries** ("productivity associated OTU's")
- **778 × 0.05 = 49 False Discoveries**

Main Error Rates

1 Family-Wise Error Rate:

$$FWER = \Pr(\# \text{False Discoveries} \geq 1)$$

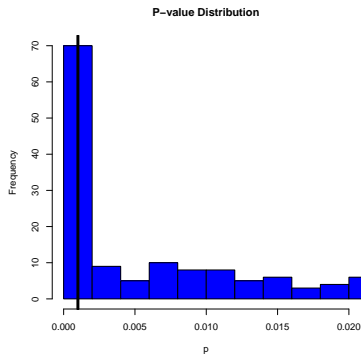
- Bonferroni: **p-value** $\leq \frac{0.05}{778} \rightarrow$ **54 discoveries**

2 False Discovery Rate:

$$FDR = E \left[\frac{\# \text{ False Discoveries}}{\# \text{ Discoveries}} \right]$$

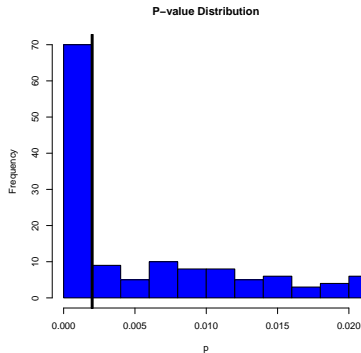
- BH procedure: **p-value** $\leq 0.005 \rightarrow$ **82 discoveries**
 - Benjamini and Hochberg (1995)

Illustration of BH procedure



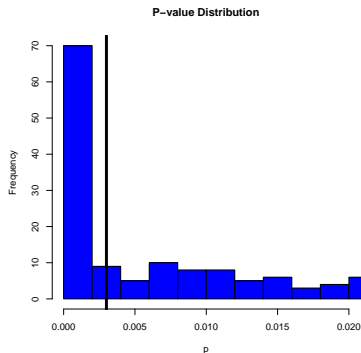
$$\widehat{FDR}(0.001) = \frac{778 \times 0.001}{54} = 0.014$$

Illustration of BH procedure



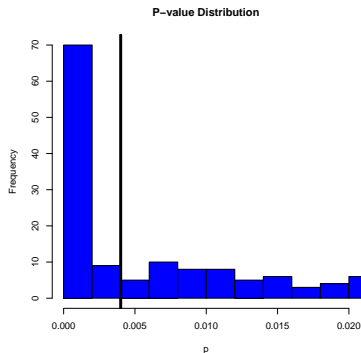
$$\widehat{\text{FDR}}(0.002) = \frac{778 \times 0.002}{64} = 0.024$$

Illustration of BH procedure



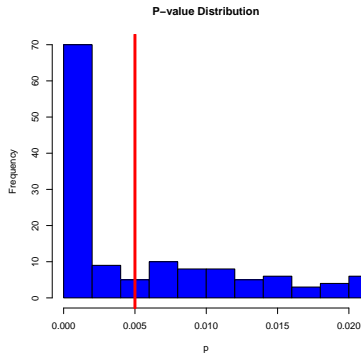
$$\widehat{\text{FDR}}(0.003) = \frac{778 \times 0.003}{70} = 0.033$$

Illustration of BH procedure



$$\widehat{FDR}(0.004) = \frac{778 \times 0.004}{74} = 0.042$$

Illustration of BH procedure



$$\widehat{FDR}(0.005) = \frac{778 \times 0.005}{82} \approx 0.05$$

Properties of BH Procedure

Theorem (Benjamini and Hochberg; 1995)

If P -values from true null hypotheses are

- 1 *independent*
- 2 *uniformly distributed*

then the BH procedure has $FDR \leq a_0 \alpha \leq \alpha$

- a_0 = proportion of true null hypotheses

FDR Areas

FDR Research Areas

Uniform Distribution

P-values **not uniformly distributed under null** if

1 Model misspecified

- Efron (2001, 2007); Pollard and van der Laan(2004)
- Habiger and Peña (2011)

2 Data/test statistic discrete

- Lancaster (1961); Pratt(1961)
- Geyer and Meeden (2005); Kulinskaya and Lewin (2009)
- Gutman and Hochberg (2007)
- Habiger (2013)

Dependence

P -values may not be **independent under null**

1 Positive Dependence

- Benjamini and Yekuteuli (2001)
- Guo, Li, and Sarkar (2013), Guo and Sarkar(2013)

2 Weak Dependence

- Storey (2001), Genovese and Wasserman (2002), Storey, Taylor and Siegmund (2004)

3 Arbitrary/Strong Dependence

- Benjaminin and Yekuteuli (2001)
- Efron (2012); Desai and Storey (2012); Fan, Han and Gu (2012)

Exhausting the α

BH: $FDR = a_0\alpha < \alpha$

1 Exact **adaptive** FDR control via \hat{a}_0

- Storey, Taylor and Siegmund (2004); Benjamini, Krieger and Yekutieli (2006); Gavrilov, Benjamini and Sarkar (2009); Liang and Nettleton (2012)

2 Consistent estimation of a_0

- Sun and Cai (2007); Jin and Cai (2007); . . . MANY more

3 Asymptotically Optimal Rejection Curve

- Finner et. al (2009)

Heterogeneity

Data **not homogeneous** - Example ancillary statistics

- 1 **weighted** BH type procedures
 - Genovese and Wasserman (2006); Roeder and Wasserman (2009); Roquain and van de Wiel (2009); Peña, Habiger and Wu (2011)
- 2 **focus on group/cluster structure**
 - Sun and Cai (2009); Hu, Zhao and Zhou (2010)

P-value efficiency

- local FDR for z -scores: $lfdr(z) = \frac{pf_0(z)}{f(z)}$
 - Efron et. al (2001); Efron(2010); Sun and Cai (2007); Jin and Cai (2007)
 - Rubin, Dudoit and van der Laan (2006); Habiger and Peña (2013)
- P-value approach = Z-value approach if p -value appropriately defined
 - Habiger (2012)

Idea of this Paper

Weak Dependence + Weighted + Adaptive

Finite FDR Control

Preliminaries and Finite FDR Control

Data and Hypotheses

- **Data:** $\mathbf{Z} = (Z_m, m \in \mathcal{M}) \sim F$ for $\mathcal{M} = \{1, 2, \dots, M\}$
 - Example: $\mathbf{Z} \sim MVN(\boldsymbol{\mu}, \boldsymbol{\Sigma})$
- **Null hypotheses:** $\mathbf{H} = (H_m, m \in \mathcal{M})$
 - Example: $H_m : \mu_m = 0$
- **True nulls:** $\mathcal{M}_0 = \{m \in \mathcal{M} : H_m \text{ true}\} \subseteq \mathcal{M}$
- **Number true nulls:** $M_0 = |\mathcal{M}_0|$

Decision functions

Decision fxn: $\delta_m(\mathbf{Z}_m; t_m) \in \{0, 1\}$ - t_m a “threshold” (or size or index)

- Example 1: $\delta_m(\mathbf{Z}_m; t_m) = I(\mathbf{Z}_m \geq \bar{\Phi}^{-1}(t_m))$
- Example 2: p -value $\rightarrow \delta_m(P_m(\mathbf{Z}_m); t_m) = I(P_m(\mathbf{Z}_m) \leq t_m)$
- Assumptions
 - 1 $E[\delta_m(\mathbf{Z}_m; t_m)] = t_m$ under H_m
 - 2 $t_m \mapsto E[\delta_m(\mathbf{Z}_m; t_m)]$ continuous and strictly increasing
- Shorthand: $\delta_m(t_m)$ a Bernoulli process

Multiple decision fxn: $\delta(\mathbf{t}) = [\delta_m(t_m), m \in \mathcal{M}]$

- $\mathbf{t} = (t_m, m \in \mathcal{M})$ - “threshold vector”

False Discovery Proportion/Rate

False Discovery Proportion (FDP):

$$FDP(\mathbf{t}) = \frac{V(\mathbf{t})}{R(\mathbf{t}) \vee 1}$$

False Discovery Rate (FDR):

$$FDR(\mathbf{t}) = E[FDP(\mathbf{t})]$$

- $V(\mathbf{t}) = \sum_{m \in \mathcal{M}_0} \delta_m(\mathbf{t}_m)$ number false discoveries
- $R(\mathbf{t}) = \sum_{m \in \mathcal{M}} \delta_m(\mathbf{t}_m)$ number discoveries

Weights and a single threshold

Goal: Choose t large as possible s.t. $FDR(t) \leq \alpha$

$$\Downarrow$$
$$t_m = \bar{t} \left(\frac{t_m}{\bar{t}} \right) \equiv t w_m$$

↙ ↘

choose t choose w

Procedure: Fixed weights

- 1 Choose threshold

$$t_\alpha^\lambda = \sup\{0 \leq t \leq \lambda : \widehat{FDR}^\lambda(t\mathbf{w}) \leq \alpha\}$$

where

$$\widehat{FDR}^\lambda(t\mathbf{w}) = \frac{\hat{M}_0(\lambda\mathbf{w})t}{R(t\mathbf{w}) \vee 1}$$

- 2 Compute $\delta(t_\alpha^\lambda \mathbf{w})$
 - Weighted **adaptive** multiple decision function (WAMDF)

Details

$$\hat{M}_0(\lambda \mathbf{w}) = \frac{M - R(\lambda \mathbf{w}) + 1}{1 - \lambda}$$

- Weighted version of Storey et. al (2004) estimator
- Intuition for $\mathbf{w} = \mathbf{1}$ and $\lambda = 1/2$:
 - If all alternative p -values near 0 then

$$\hat{M}_0 = \{\#p\text{-values} > 1/2\} \times 2$$

- Should have $\lambda w_m < 1$

FDR bound

Lemma (FDR bound)

For $M_0 \geq 1$ and under

(A1) $(Z_m, m \in \mathcal{M}_0)$ independent collection

(A2) $\lambda w_m < 1$ for every m ,

$$FDR(t_\alpha^\lambda \mathbf{w}) \leq \alpha \bar{w}_0 \frac{1 - \lambda}{1 - \lambda \bar{w}_0}$$

- $\bar{w}_0 = M_0^{-1} \sum_{m \in \mathcal{M}_0} w_m$ average null weight
- Corollary: $\mathbf{w} = \mathbf{1} \Rightarrow \bar{w}_0 = 1 \Rightarrow$ Theorem 3 - Storey et al (2004)

FDR Control

Problem: \bar{w}_0 unobservable!

Theorem (FDR control)

Let $w_{(M)} = \max\{w_m, m \in \mathcal{M}\}$ and define

$$\alpha^* = \alpha \frac{1}{w_{(M)}} \frac{1 - \lambda w_{(M)}}{1 - \lambda}.$$

Then under (A1) and (A2), $FDR(t_{\alpha^*}^\lambda \mathbf{w}) \leq \alpha$.

- Generally have $\bar{w}_0 \leq 1 \dots$

Asymptotic results

Asymptotic FDP Control

- (Weighted) adaptive vs. unadaptive vs. Oracle
- “ α -exhaustion”

Remark: Notation - $w_{0,M}, \widehat{FDR}_M, \dots$

Assumptions

- **Weak Dependence:** for $0 \leq t \leq u$,

(A3) $R(t\mathbf{w}_M)/M \rightarrow G(t)$ a. s.

(A4) $V(t\mathbf{w}_M)/M \rightarrow a_0\mu_0 t$ a. s.

- $\bar{w}_{0,M} \rightarrow \mu_0$ asymptotic mean null weight
- $\frac{M_0}{M} \rightarrow a_0$ asymptotic proportion true nulls
- $u = \sup\{t : tw_m \leq 1\}$

- **FDR controllable:**

(A5) $t/G(t)$ is strictly increasing and continuous with

$$\lim_{t \downarrow 0} \frac{t}{G(t)} = 0 \quad \text{and} \quad \lim_{t \uparrow u} \frac{t}{G(t)} = \frac{u}{G(u)} \leq 1,$$

- Similar to Genovese and Wasserman (2006), Storey et. al(2004), ...

FDR Estimators

Estimators + limits

Unadaptive $\widehat{FDR}_M^0(t\mathbf{w}_M) = \frac{Mt}{R(t\mathbf{w}_M)} \rightarrow \frac{t}{G(t)} \equiv \widehat{FDR}_\infty^0(t)$

Adaptive $\widehat{FDR}_M^\lambda(t\mathbf{w}_M) = \frac{\hat{M}_0(\lambda\mathbf{w}_M)t}{R(t\mathbf{w}_M)} \rightarrow \frac{1-G(\lambda)}{1-\lambda} \frac{t}{G(t)} \equiv \widehat{FDR}_\infty^\lambda(t)$

Oracle $FDP_M(t\mathbf{w}_M) = \frac{V(t\mathbf{w}_M)}{R(t\mathbf{w}_M)} \rightarrow \frac{a_0\mu_0 t}{G(t)} \equiv FDP_\infty(t)$

Lemma (Glivenko-Cantelli)

Under (A2) - (A5), convergence is uniform for each “estimator” a.s., i.e.

$$\sup_{0 \leq t \leq u} |\widehat{FDR}_M(t\mathbf{w}) - \widehat{FDR}_\infty(t)| \rightarrow 0 \quad \text{almost surely}$$

Thresholds

Example: $t_{\alpha, M}^{\lambda} = \sup\{0 \leq t \leq u : \widehat{FDR}_M^{\lambda}(t\mathbf{w}) \leq \alpha\}$

	Threshold	
	Finite (M) (random)	Asymptotic (∞) (not random)
Unadaptive - \widehat{FDR}^0	$t_{\alpha, M}^0$	$t_{\alpha, \infty}^0$
Adaptive - \widehat{FDR}^{λ}	$t_{\alpha, M}^{\lambda}$	$t_{\alpha, \infty}^{\lambda}$
Oracle FDP	$t_{\alpha, M}$	$t_{\alpha, \infty}$

Threshold comparison

Theorem (Thresholds converge)

Consider

$$\lim_{M \rightarrow \infty} t_{\alpha, M}^0 = t_{\alpha, \infty}^0 \leq \lim_{M \rightarrow \infty} t_{\alpha, M}^\lambda = t_{\alpha, \infty}^\lambda \leq \lim_{M \rightarrow \infty} t_{\alpha, M} = t_{\alpha, \infty}$$

Under (A2) - (A5),

- all equalities + first inequality satisfied a.s.
 - last inequality satisfied a.s. if $\mu_0 \leq 1$.
-
- Thresholds constant
 - **Adaptive threshold** > unadaptive threshold

FDP comparison

Theorem (FDPs converge)

$$\lim_{M \rightarrow \infty} \text{FDP}_M(t_{\alpha, M}^0 \mathbf{w}_M) \leq \lim_{M \rightarrow \infty} \text{FDP}_M(t_{\alpha, M}^\lambda \mathbf{w}_M) \leq \lim_{M \rightarrow \infty} \text{FDP}_M(t_{\alpha, M} \mathbf{w}_M) = \alpha$$

Under (A2) - (A5),

- *first inequality and last equality satisfied a.s.*
- *last inequality satisfied a.s. if $\mu_0 \leq 1$*
- **Important: FDP \neq FDR - Error control on 1 replication**
- **Q: Does $\text{FDP}_M(t_{\alpha, M}^\lambda \mathbf{w}_M) \rightarrow \alpha$?**

α -exhaustion idea

“Good procedures should be *α -exhaustive*” - Have $FDR_M \rightarrow \alpha$ under some “least favorable distribution”

- Finner et. al (2009); Roquain and Villers (2011) - AORC:

$$p_{(k)} \leq \alpha \frac{k}{M - k(1 - \alpha)}$$

- **Least favorable** dist under **i.i.d.** is “Dirac-Uniform (DU)”

$$E[\delta_m(t)] = \begin{cases} t & m \in \mathcal{M}_0 \\ 1 & m \in \mathcal{M}_1 \end{cases}$$

- **Step-up procedures fail**

α -exhaustion

Theorem (α -exhaustion)

If $\mu_0 = 1$, then under weak dependence (A3 - A4) and a DU distribution, $FDP_M(t_{\alpha, M}^\lambda \mathbf{w}_M) \rightarrow \alpha$ almost surely, i.e. $\delta(t_{\alpha, M}^\lambda \mathbf{w}_M)$ is α -exhaustive.

Corollary (Adaptive BH is α -exhaustive)

The (unweighted) adaptive BH procedure **step up** procedure - Storey et. al (2004)- is " α -exhaustive" under weak dependence (A3 - A4).

- Extends Finner et. al (2009) theory to 1) **weighted** 2) **adaptive** 3) **step-up** procedure under 4) **weak dependence**

Question

Questions

- 1 α -exhaustion \Rightarrow “optimal”
- 2 How to choose \mathbf{w}

Optimal Weights

Optimal Weights

- Mixture model
- Optimal weights for fixed t
- Optimal weights for approximation of $t_{\alpha, \infty}^{\lambda}$

Mixture Model

Model (Random effects model)

Let $(Z_m, \theta_m, p_m, \gamma_m)$, $m \in \mathcal{M}$ be i.i.d. random vectors with

$$F(z_m | \theta_m, \gamma_m) = (1 - \theta_m)F_0(z_m) + \theta_m F_1(z_m | \gamma_m)$$

and

$$F(z_m | p_m, \gamma_m) = (1 - p_m)F_0(z_m) + p_m F_1(z_m | \gamma_m).$$

- $\theta_m \in \{0, 1\}$ with $p_m = \Pr(\theta_m = 1)$ ind. of γ_m
- $E[p_m] = 1 - a_0$ for $0 < a_0 < 1$.
- $F_0(\cdot)$ “null distribution” and $F_1(\cdot | \gamma_m)$ “alternative distribution”
- **Heterogeneity: effect size γ_m and prior probability p_m**

Power function

For $t_m = tw_m$,

$$\pi_{\gamma_m}(t_m) = E[\delta_m(Z_m; t_m) | \theta_m = 1, \gamma_m]$$

is the **power function**

(A6) $t_m \mapsto \pi_{\gamma_m}(t_m)$ is concave and twice differentiable for $t_m \in (0, 1)$ with $\lim_{t_m \uparrow 1} \pi'_{\gamma_m}(t_m) = 0$ and $\lim_{t_m \downarrow 0} \pi'_{\gamma_m}(t_m) = \infty$ a.s.

- Similar to Genovese and Wasserman (2006); Peña, Habiger, Wu (2011), ...

Optimality goal

- 1 Assume $\bar{t} = t$ fixed \Leftrightarrow (ex. Bonferroni $t = \alpha/M$)
- 2 Goal: Maximize ETP / power

$$\pi(\mathbf{t}, \mathbf{p}, \gamma) \equiv E \left[\sum_{m \in \mathcal{M}} \theta_m \delta_m(t_m) \mid \gamma, \mathbf{p} \right] = \sum_{m \in \mathcal{M}} p_m \pi_{\gamma_m}(t_m),$$

subject to $\bar{t} = t$.

- 3 Can always recover weights

$$t_m = t w_m \Rightarrow w_m = \frac{t_m}{t}$$

Optimal fixed- t threshold/weights

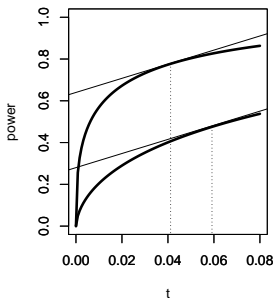
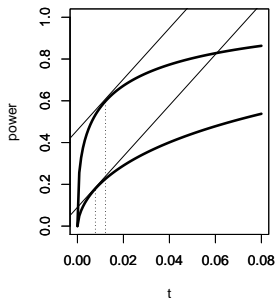
Theorem (Optimal Fixed- t threshold)

Under (A6) and the random effects model, for any fixed $t \in (0, 1)$, the maximum of $\pi(t, \mathbf{p}, \gamma)$ with respect to t subject to constraint $\bar{t} = t$ exists, is unique, and satisfies (a.s.)

$$\pi'_{\gamma_m}(t_m) = k/p_m$$

- 1 For any k can compute $t_m(k/p_m, \gamma_m)$
- 2 Find the k^* satisfying constraint
- 3 Compute optimal fixed- t weights $w_m^* = \frac{t_m(k^*/p_m, \gamma_m)}{t}$

Illustration: $p_1 = p_2 = 1/2$



- $t = 0.01, 0.05 \Rightarrow k^* = 6.1, 1.7$

Approximation Idea

Problem: $t_{\alpha, M}^{\lambda}$ not fixed so previous theorem not applicable

Solution: Recall $t_{\alpha, M}^{\lambda} \rightarrow t_{\alpha, \infty}^{\lambda}$

- Can approximate $t_{\alpha, \infty}^{\lambda}$ “well” using \mathbf{p} and γ (details omitted)

Assessment

Assessment

Notation

- **Asymptotically optimal** fixed- t weights \mathbf{w}_M^* (using $t_{\alpha, \infty}^\lambda$)
- **Approximately optimal** fixed- t weights $\hat{\mathbf{w}}_M$ (using approximation $t_{\alpha, \infty}^\lambda$)
- **Perturbed** fixed- t weights $\tilde{\mathbf{w}}_M$ ($\tilde{w}_{m, M} = U_m \hat{w}_{m, M}$)

FDP control

Theorem (FDP control)

Under the Random Effects Model and (A6), conditions (A2) - (A5) are satisfied and $\mu_0 \leq 1$. Hence, almost surely

$$\lim_{M \rightarrow \infty} FDP_M(t_{\alpha, M}^0 \tilde{\mathbf{w}}_M) \leq \lim_{M \rightarrow \infty} FDP_M(t_{\alpha, M}^\lambda \tilde{\mathbf{w}}_M) \leq \alpha$$

For large M

- **adaptive weighted procedure** DOMINATES **unadaptive weighted procedure** and is **valid** even if **weights are misspecified**

α -exhaustion

Theorem (α -exhaustion)

Under model 1 and (A6), $\delta(t_{\alpha, M} \tilde{\mathbf{w}}_M)$ is α -exhaustive if $p_1 = p_2 = \dots = p_M = p$

- 1 Many optimal weighting schemes can be motivated using random effects model with $p_1 = p_2 = \dots = p_M$
 - Spjotvoll (1972), Genovese and Wasserman (2006); Storey(2007); Peña, Habiger, Wu (2011)
- 2 If used with $t_{\alpha, M}^\lambda \rightarrow$ Optimally weighted + α -exhaustive

Optimal Weights

Theorem (Asymptotically optimal weights)

Under Model 1 and (A6), $\delta_m(t_{\alpha, M}^\lambda \hat{w}_m) \rightarrow \delta_m(t_{\alpha, \infty}^\lambda w_m^)$ almost surely for every m .*

- Approximately optimal weights are asymptotically optimal

Simulation setup

Data: $Z_m \sim N(\theta_m \gamma_m, 1)$

Decision: $\delta_m(Z_m; t_m) = I(Z_m \geq \bar{\Phi}^{-1}(t_m))$

Effect sizes: $\gamma_m \stackrel{i.i.d.}{\sim} Un(1, 1), Un(1, 3), \text{ or } Un(1, 5)$

Procedures:

UU ($\mathbf{1}, t_{\alpha, M}^0$) unweighted + unadaptive - Benjamini and Hochber (1995)

WU ($\mathbf{w}_M^*, t_{\alpha, M}^0$) weighted + unadaptive - Genovese and Wasserman (2006);
Peña, Habiger, Wu (2011)

UA ($\mathbf{1}, t_{\alpha, M}^\lambda$) = unweighted + adaptive - Storey et. al (2004)

WA ($\mathbf{w}_M^*, t_{\alpha, M}^\lambda$) **weighted + adaptive** - Habiger (201?)

Some Heterogeneity

Simulation 1

	$\gamma_m = 1$	$\gamma_m \stackrel{i.i.d.}{\sim} Un(1, 3)$	$\gamma_m \stackrel{i.i.d.}{\sim} Un(1, 5)$
UU	0.007(0.025)	0.390(0.024)	0.707(0.025)
WU	0.007(0.025)	0.395(0.025)	0.729(0.025)
UA	0.009 (0.030)	0.454 (0.034)	0.753(0.039)
WA	0.009 (0.030)	0.457 (0.035)	0.778 (0.039)

Average Power (FDR) for $p_m = 1/2$

- WA - Optimally Weighted + α -exhaustive
- UA - α -exhaustive

More Heterogeneity

Simulation 2

	$\gamma_m = 1$	$\gamma_m \stackrel{i.i.d.}{\sim} Un(1, 3)$	$\gamma_m \stackrel{i.i.d.}{\sim} Un(1, 5)$
UU	0.007(0.025)	0.390(0.025)	0.711(0.025)
WU	0.012(0.012)	0.431(0.015)	0.755(0.016)
UA	0.009(0.026)	0.457(0.035)	0.757(0.039)
WA	0.017 (0.015)	0.504 (0.021)	0.807 (0.026)

Average Power (FDR) for $p_m \stackrel{i.i.d.}{\sim} Un(0, 1)$

- WA Optimally weighted
- UA - α -exhaustive

Perturbed weights

Simulation 3

	$\gamma_m = 1$	$\gamma_m \stackrel{i.i.d.}{\sim} Un(1, 3)$	$\gamma_m \stackrel{i.i.d.}{\sim} Un(1, 5)$
UU	0.006(0.025)	0.391(0.026)	0.709(0.025)
WU	0.012(0.013)	0.394(0.016)	0.724(0.016)
UA	0.008(0.028)	0.458(0.036)	0.756(0.039)
WA	0.019(0.015)	0.457(0.022)	0.773(0.026)

Average Power (FDR) for $p_m \stackrel{i.i.d.}{\sim} Un(0, 1)$ and perturbed weights

- UA - α -exhaustive

Concluding Remarks

Concluding Remarks

Summary of results

- Optimality of weighted adaptive procedure
 - Heterogeneity **and** adaptive vs. Heterogeneity **or** adaptive
- Robustness of weighted adaptive procedure
 - FDP (not FDR) control under weak dependence and noisy weights
 - Good power under noisy weights

Near future work

OTU#	Sufficient Stat	Ancillary Stat	weights
1	18.14	7	?
2	19.13	14	?
⋮	⋮	⋮	⋮
M = 778	161.05	87	?

- How can we estimate $\gamma_m s / p_m s$?
- Different optimality goal: maximize $\sum_m \sum_j (\hat{y}_{mj} - \bar{y}_{m\cdot})^2$
 s.t. 5% of “declared productivity-associated variability
 is falsely declared variability”

Near future work continued

- 1 Choice of λ ?
 - Dynamic - Liang and Nettleton (2012)
 - Variance vs. bias vs. power
- 2 Estimation of μ_0 ?

General Future work

Optimally weighted
+
 α - exhaustive
+
strong dependence
+
discrete data

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