# ANOVA on Ranks with Bounded Influence Function

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## **Presentation Topics**

- 1. Introduction to Anova on Ranks.
  - (a) Rank Statistics
  - (b) Analysis of Variance on Ranks
  - (c) Robust Statistics & Why We Need it
  - (d) Bounded Influence ANOVA on Ranks
- 2. Application Example on MK-0869 Data.
- 3. Simulation Study.
- 4. Concluding Remarks.

## **Rank Statistics**

- Let  $\mathbf{R}^* = (R_1^*, \dots, R_N^*)$  be the vector of ranks, then  $\mathbf{R}^*$  is uniformly distributed over all permutations of the integers  $1, \dots, N$ , with density  $f(\mathbf{R}^*) = \frac{1}{N!}$ .
- Let  $V(\mathbf{R}^*)$  be a sample statistics only through  $\mathbf{R}^*$ , then  $V(\mathbf{R}^*)$  is distribution free from the raw data.  $V(\mathbf{R}^*)$  is called **rank statistics**
- Example rank statistics: for two-samples location problem with sample sizes m and n, there are Wilcoxon statistics  $W = \sum_{i=1}^{n} R_i$ , and Mann-Whitney statistics  $U = \sum_{i=1}^{m} \sum_{j=1}^{n} \Psi(Y_j X_i)$ ,  $\Psi(t) = 1$ , 0 as  $t \ge 0$ .
- Under the null hypothesis  $H_o$ :  $\Delta = 0$ , W is distributed as geometric distribution, regardless the raw data distribution.
- For large sample sizes, the Mann-Whitney-Wilcoxon statistics can be approximated by standard normal distribution.

## Example:

For two samples (X, Y) with sizes m = 3 and n = 2, there are  $\binom{3+2}{2} = 10$  arrangements to be examined to have the null distribution of W.

Arrangement	Value of $\it W$
xxxyy	W = 9
xxyxy	W = 8
xyxxy	W = 7
yxxxy	W = 6
xxyyx	W = 7
xyxyx	W = 6
yxxyx	W = 5
xyyxx	W = 5
yxyxx	W = 4
yyxxx	W = 3

#### To test:

 $H_o: \mu_1 - \mu_2 = 0 \text{ vs}$  $H_a: \mu_1 - \mu_2 > 0.$ 

Decision rule: reject  $H_o$  if  $W > w(\alpha, m, n)$ .  $w(\alpha, m, n)$  is the upper 100 $\alpha$ % percentile of sample W values.

Sometimes people referred this kind of test based on permutation theory to as "<u>Exact Test</u>" (or "Exact P-vale", "Distribution Free Test").

#### **ANOVA** on Ranks

#### <u>Friedman ranks:</u>

For each j (j=1, ..., b), let  $R_{ij}$  be the rank of  $X_{ij}$  among the observations  $X_{1j}, \ldots, x_{tj}$  within block j. The  $\{R_{ij}\}$  will be referred to as Friedman ranks.

#### Rank transform:

Let  $Q_{ij}$  be the rank of  $X_{ij}$  among all tb observations  $X_{1j}, \ldots, x_{tj}$ . "Rank transform" refers to replacing the  $X_{ij}$  with the pooled-data ranks  $\{Q_{ij}\}$ .

## Aligned ranks:

Let  $\hat{\theta}_j$  be an estimator of location for block j, and let  $A_{ij}$  be the rank of  $(X_{ij} - \hat{\theta}_j)$  among the pooled se of centered observations  $(X_{ij} - \hat{\theta}_j), \ldots, (X_{ij} - \hat{\theta}_j)$ . Then  $\{A_{ij}\}$  are called aligned ranks (Lehmann, 1975, §6.3).

#### Friedman's statistic

$$T = \frac{12b}{t(t+1)} \sum_{i=1}^{t} \{\bar{R}_{i.} - \frac{1}{2}(t+1)\}^{2}$$

under  $H_o$ , as  $b \to \infty$ ,  $T \stackrel{dist}{\to} \chi^2_{t-1}$ , where t and b are the number of treatments and patients in on-way mixed model.

#### Alternative to Friedman's statistic

Substitute actual data by Friedman's rank, do regular ANOVA basing critical values on the F(t-1,(t-1)(b-1)) distribution. This is often referred to as "ANOVA on Ranks".

## **Robust Statistics**

Statistical techniques diminishing the effect of the unusual cases is called **robust statistics**. In general  $\hat{\mu}$  is obtained by minimizing

$$\sum_{i=1}^{n} \rho(y_i - \mu)$$

or

$$\sum_{i=1}^n \psi(y_i - \widehat{\mu}) = 0$$

where  $\psi(t) = \rho'(t)$ .

1. <u>M-Estimators</u>: need to minimize  $\sum_{i=1}^n \rho(\frac{y_i - \mathbf{x}_i b}{s}) + n \log s$ , then obtain the MLE b of  $\beta$  solves  $\sum_{i=1}^n \mathbf{x}_i \psi(\frac{y_i - \mathbf{x}_i b}{s}) = 0$ 

- 2. <u>L<sub>1</sub>-Estimators</u>: Find  $\beta$  of b to minimize  $\sum_{i=1}^{n} |y_i \mathbf{x}_i b|$
- 3. Bounded Influence Functions.

# Why Robust Statistics?

- 1. Users, even expert statisticians, do not always screen the data.
- 2. The sharp decision to keep or reject an observation is wasteful. We can do better by downweighting dubious obserations than by rejecting them, although we may wish to reject completely wrong observations.
- 3. It can be difficult or even impossible to spot outliers in multivatriate or highly structured data.
- 4. Rejecting outliers affects the distribution theory, which ought to be adjusted. In particular, variances will be under-estimated from the 'cleaned' data.

#### Application Example

## Prevention of Acute and Delayed Chemotheray-Induced Emesis Associated With High-Dose Cisplatin (MK-0869)

## Study Design

	Day 1	2	3	4	5
Standard					
Low dose		VAS,	# of Emesis		
High dose					

## Objective of analysis

Do, in terms of severity measured VAS, the patients taking MK-0869 have statistically significant lower nausea symptom?

#### Variables

- Number of emesis per day.
- Daily <u>n</u>ausea visual <u>a</u>nalogue <u>s</u>cale (VAS) score (as a measure of severity of nausea)
- Other adverse experiences, demographics, etc..

## MK-0869 data

$$X_{ijk} = \mu + \alpha_i + \gamma_j + \epsilon_{ijk}$$
 
$$i = 1, 2, \text{3 treatments;} \quad j = 1, \dots, 151 \text{ patients.}$$
 
$$k = 1, \dots, 5 \text{ day into medicine;}$$
 
$$\epsilon_{ijk} \overset{dist}{\sim} \text{ any distribution, } X = \text{VAS}$$

# Treatment Effects Significance Table ( $\alpha = 0.05\%$ )

		Bounding with	Function without
Friedman ranks	F-value	123.47	46.99
	Pr(F)	<b>0</b>	<b>0</b>
Rank transform	F-value	3.539	2.408
	Pr(F)	<b>0.0295</b>	0.0907
Aligned ranks	F-value	133.672	3.547
	Pr(F)	<b>0</b>	<b>0.0293</b>

# **Limited Simulation Study Result**

$$n_1 = 60, n_2 = 60, n_3 = 30, \alpha = 0.05$$
# repeat = 500

## • Rank Transform

	Percentage Significant	
Treatment Effect	with	without
None	3.5%	5.1%
Low	65.2%	58.9%
High	65.9 %	61.8%

## Friedman Ranks

	Percentage Significant	
Treatment Effect	with	without
None	4.5%	4.9%
Low	4.5% 51.5%	50.2%
High	65.2%	66.2%

# Aligned Ranks

	Percentage Significant	
Treatment Effect	with	without
None	5.5%	4.7%
Low	50.1%	55.5%
High	67.5%	60.4%

## **Concluding Remarks**

- The power of ANOVA tests is reduced with long-tailed data, as proved in the classical theory.
- ANOVA on ranks procedure is distribution free, and in general will be superior to standard ANOVA for data with frequent extreme values.
- But different ranking schemes give different Pvalues, due to the pattern of extreme observations.
- A pre-defined bounding influence function based on available information unifies 3 main types of ranks, and <u>seemingly</u> increases the robustness of *P*-value as well as the power of the test in <u>limited</u> simulation study.
- ANOVA on ranks theory is validated for fixed t as  $b \to \infty$  situation. When b is small, only empirical results available.

#### References

#### ANOVA on ranks

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