Lecture 40 Nonparametric Methods

BIO210 Biostatistics

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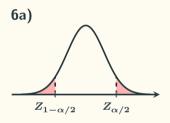
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Parametric Tests

- 1. Specify what you are comparing.
- 2. Formulate hypotheses
- 3. Check assumptions
- 4. Determine significance level α
- 5. Compute the test statistic
- 6. Check significance
- 7. Make a decision about whether to reject H_0
- 8. Interpret findings



- **6b)** Calculate the p-value.
- **6c)** Construct $(1 \alpha) \times 100\%$ confidence interval to see if it covers the H_0 value.

Nonparametric Methods

Nonparametric tests do not rely on data following specific distribution (e.g. normal). They are also called distribution-free methods and are often used when the assumptions of parametric tests are violated.

- Wilcoxon, Frank (1945) Individual comparisons by ranking methods. Biometrics Bulletin. 1 (6): 80–83.
- Wilcoxon Sign Test
- Wilcoxon Signed-Rank Test
- Wilcoxon Rank Sum Test (Mann-Whitney U Test)

Nonparametric Test 1 - Wilcoxon Sign Test

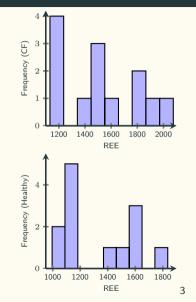
Wilcoxon Sign Test:

A test for median of paired data.

Compute the difference Record the sign Remove zeros and update sample size Record the number of positive signs (D)Use D to calculate the p-value

Resting energy expenditure (REE) for patients with cystic fibrosis and healthy individuals matched on age, sex, height and weight.

| Pair | REE (CF | kcal/day) Healthy | Difference | Sign |
|------|-------------|----------------------|------------|------|
| 1 | 1153 | 996 | 157 | + |
| 2 | 1132 | 1080 | 52 | + |
| 3 | 1165 | 1182 | -17 | - |
| 4 | 1460 | 1452 | 8 | + |
| 5 | 1634 | 1162 | 472 | + |
| 6 | 1493 | 1619 | -126 | - |
| 7 | 1358 | 1140 | 218 | + |
| 8 | 1453 | 1123 | 330 | + |
| 9 | 1185 | 1113 | 72 | + |
| 10 | 1824 | 1463 | 361 | + |
| 11 | 1793 | 1632 | 161 | + |
| 12 | 1930 | 1614 | 316 | + |
| 13 | 2075 | 1836 | 239 | + |



Nonparametric Test 1 - Wilcoxon Sign Test

- 1. The null/alternative hypotheses:
- H_0 : no difference in REE between CF and healthy people
- H_1 : there is a difference
- 2. If H_0 were true, we should expect similar number of "+" and "-" signs.
- 3. Under H_0 :

$$D \sim B(n,0.5) \begin{cases} D \sim \mathcal{N}\left(\frac{n}{2},\,\frac{n}{4}\right), & \text{if the sample size is large enough} \\ & (i.e. \text{ both } D \text{ and } n-D \text{ are more than 10}) \end{cases}$$

$$D \sim B(n,0.5), & \text{otherwise}$$

Nonparametric Test 2 - Wilcoxon Signed-Rank Test

Wilcoxon Signed-Rank Test: a test for median of paired data, but taking into account the magnitude of the difference.

| the magnitude of the difference. | | | | | | | | | | |
|---------------------------------------|--|--|--|--|--|--|--|--|--|--|
| Compute the difference | | | | | | | | | | |
| \ | | | | | | | | | | |
| Rank the difference by the absolute | | | | | | | | | | |
| values (the smallest to the largest) | | | | | | | | | | |
| \ | | | | | | | | | | |
| Update the sign of the rank | | | | | | | | | | |
| by the sign of the difference | | | | | | | | | | |
| • | | | | | | | | | | |
| Sum the positive and negative ranks, | | | | | | | | | | |
| separately and let T be the smaller | | | | | | | | | | |
| sum of rank, ignoring the sign | | | | | | | | | | |
| · | | | | | | | | | | |

Use T to calculate the p-value

| Pair | REE (| (kcal/day) Healthy | | Rank | Sign + | ed rank - |
|------|-------|-----------------------|------|------|-----------|--------------|
| 1 | 1153 | 996 | 157 | 6 | 6 | |
| 2 | 1132 | 1080 | 52 | 3 | 3 | |
| 3 | 1165 | 1182 | -17 | 2 | | -2 |
| 4 | 1460 | 1452 | 8 | 1 | 1 | |
| 5 | 1634 | 1162 | 472 | 13 | 13 | |
| 6 | 1493 | 1619 | -126 | 5 | | -5 |
| 7 | 1358 | 1140 | 218 | 8 | 8 | |
| 8 | 1453 | 1123 | 330 | 11 | 11 | |
| 9 | 1185 | 1113 | 72 | 4 | 4 | |
| 10 | 1824 | 1463 | 361 | 12 | 12 | |
| 11 | 1793 | 1632 | 161 | 7 | 7 | |
| 12 | 1930 | 1614 | 316 | 10 | 10 | |
| 13 | 2075 | 1836 | 239 | 9 | 9 | |
| | | | | Sum | 84 | -7 |

Nonparametric Test 2 - Wilcoxon Signed-Rank Test

- 1. The null/alternative hypotheses:
- H_0 : no difference in REE between CF and healthy people
- H_1 : there is a difference
- 2. If H_0 were true, we should expect similar number of "+" and "-" signs, and the absolute values of the sum of positive ranks and the sum of negative ranks should be comparable.
- 3. Under H_0 :

$$Z_T=\frac{T-\mu_T}{\sigma_T}\sim\mathcal{N}(0,1)$$
 where $\mu_T=\frac{n(n+1)}{4},$ and $\sigma_T=\sqrt{\frac{n(n+1)(2n+1)}{24}}$

- Wilcoxon Rank Sum Test (Mann–Whitney U Test)
- A test for median of two independent samples, taking into account the magnitude of the difference.
- Data: Normalised mental age (nMA) in two populations of children suffering from phenylketonuria (unable to metabolise phenylalanine). It has been suggested that an elevated level of serum phenylalanine increases a child's likelihood of mental deficiency.

| | Low Ex | posure | | High Exposure | | | | | | |
|------|--------|--------|------|---------------|------|------|------|--|--|--|
| 34.5 | 47.5 | 54.0 | 37.5 | 28.0 | 45.5 | 52.0 | 35.0 | | | |
| 48.7 | 54.0 | 39.5 | 49.0 | 46.0 | 53.0 | 37.0 | 48.0 | | | |
| 55.0 | 40.0 | 51.0 | 56.5 | 53.0 | 37.0 | 48.3 | 54.0 | | | |
| 45.5 | 51.0 | 57.0 | 47.0 | 43.5 | 48.7 | 54.0 | 44.0 | | | |
| 52.0 | 58.5 | 47.0 | 53.0 | 51.0 | 55.0 | | | | | |
| 58.5 | | | | | | | | | | |

Wilcoxon Rank Sum Test

- Step 1: Treat two samples as one, rank by the magnitude (smallest as rank 1) while keep tracking the source of the data.

| Data: Rank: | | | | | | | |
|----------------|--|--|--|--|--|--|--|
| Data: Rank: | | | | | | | |
| Data: Rank: | | | | | | | |

Wilcoxon Rank Sum Test

- Step 2: Identify tied values, and update the rank with the average of their ranks.

| Data: | 28.0 | 34.5 | 35.0 | 37.0 | 37.0 | 37.5 | 39.5 | 40.0 | 43.5 | 44.0 | 45.5 | 45.5 | 46.0 |
|----------------|------|------|------|------|------|------|------|------|------|------|------|------|------|
| | | | | | | | | | 9 | | | | |
| Data: Rank: | 47.0 | 47.0 | 47.5 | 48.0 | 48.3 | 48.7 | 48.7 | 49.0 | 51.0 | 51.0 | 51.0 | 52.0 | 52.0 |
| Rank: | 14.5 | 14.5 | 16 | 17 | 18 | 19.5 | 19.5 | 21 | 23 | 23 | 23 | 25.5 | 25.5 |
| Data: Rank: | 53.0 | 53.0 | 53.0 | 54.0 | 54.0 | 54.0 | 54.0 | 55.0 | 55.0 | 56.5 | 57.0 | 58.5 | 58.5 |
| Rank: | 28 | 28 | 28 | 31.5 | 31.5 | 31.5 | 31.5 | 34.5 | 34.5 | 36 | 37 | 38.5 | 38.5 |

Wilcoxon Rank Sum Test

- Step 3: Sum the ranks in each group separately, and let ${\it W}$ be the smaller sum:

Rank Sum (Low Exposure): 467 Rank Sum (High Exposure): 313
$$\Rightarrow W = 313$$

- Step 4: Compute the test statistics:

$$Z_W=\frac{W-\mu_W}{\sigma_W}$$
 where $\mu_W=\frac{n_S(n_S+n_L+1)}{2},$ and $\sigma_W=\sqrt{\frac{n_Sn_L(n_S+n_L+1)}{12}}$

 n_S : is the sample size of the group that has the smaller rank sum n_L : is the sample size of the group that has the larger rank sum

Wilcoxon Rank Sum Test

- Step 5: calculate the p-value:
 - 5.1) The null/alternative hypotheses: H_0 : no difference in nMA medians between low and high phenylalanine exposure H_1 : there is a difference
 - 5.2) If H_0 were true, we should expect the ranks to be distributed randomly between the two groups. Therefore, the average ranks for each sample should be approximately equal.
 - 5.3) Under H_0 :

$$Z_W \sim \mathcal{N}(0,1)$$

5.4) The p-value can be calculated using the above property.

Nonparametric Methods

• Advantages: fewer assumptions, population distribution don't need to be normally distributed, test statistics based on ranks are easier to calculate.

• Disadvantages: less power, need larger sample size, not considering all information (i.e. only ranks).