Test Taking Strategies for BPS Exams Conquering the Biostatistics Question

Anthony J. Busti, MD, PharmD, MSc, FNLA, FAHA

HIGH-YIELD MED REVIEWS

Disclaimer

- High-Yield Med Reviews has no working relationship with BPS (Board of Pharmacy Specialities)
- This live webinar event is not endorsed or sponsored by BPS or anyone other then High-Yield Med Reviews
- This is not meant to be a commercial or sales pitch



Introduction



Anthony Busti, MD, PharmD, MSc, FNLA, FAHA

Agenda

- A General Overview
- Part 1 Conquering the Biostatistics Question
- Part 2 Interpreting Statistical Results Correctly
- A Special Coupon Code
- Live Q&A

HIGH-YIELD MED REVIEWS



Participation Required You Must Fill in Some Blanks Tackling the Biostatistics Question
A Quick Recap – Initial Steps





Conquering the Biostatistics Question

- Essential steps and decision points
 - 1. Consider drawing out study design in question
 - 2. How many groups are being studied?
 - a. Are those groups related or independent of each other?
 - 3. What type of data is represented in the outcome of interest (i.e., nominal, ordinal, continuous)?
 - 4. Connect the row and column on summary table
 - 5. The Killer Foil Moment \rightarrow If "applicable" results/data are available, consider the following:
 - a. How many patients are in each group?
 - b. Does it appear to be parametric or nonparametric?



Tackling the Biostatistics Question Memorize this Chart



| Type of Data | Two Independent Samples | Related or Paired Samples | 3 or more Independent Samples | 3 or more Related Samples | Measures of Correlation |
|--------------|--|---|--|---------------------------------|--|
| Nominal | 1.Chi-square 2.Fisher's Exact | McNemar Test | Chi-square for k independe nt samples | Cochran Q | Contingency coefficient |
| Ordinal | 1.Mann- Whitney U 2.Wilcoxon Rank Sum | 1.Sign test 2.Wilcoxon Signed Rank | Kruskal- Wallis one way ANOVA | Freidman 2 way ANOVA | 1.Spearman 2.Kendal rank 3.Kendal Coe |
| Continuous | 1.Student's t-test 2.Mann- Whitney U | Paired t-test | 1-way ANOVA | 2-way ANOVA | Pearson's Correlation |

Tackling the Biostatistics Question Interpreting the Statistical Results Correctly



Treatment

Patients were randomly assigned to receive dexamethasone sodium phosphate (Oradoxn), at a dose of 10 mg given every six hours intravenously for four days, or placebo that was identical in appearance to the active drug. The study medication was given 15 to 20 minutes before the parenteral administration of antibiotics. After the interim analysis, the protocol was amended to allow adminis-tration of the study medication with the antibiotics.

Balanced treatment assignments within each hospital were achieved with the use of a computer-generated list of random numbers in blocks of six. The code was not broken until the last patient to be enrolled had completed eight weeks of follow-up. Treatment

The New England Journal of Medicine Copyright © 2002 by the Mar etts Medical S

NOVEMBER 14, 2002 VOLUME 347 DEXAMETHASONE IN ADULTS WITH BACTERIAL MENINGITIS

JAN DE GANS, PH.D., AND DIEDERIK VAN DE BEEK, M.D., FOR THE EUROPEAN DEXAMETHASONE IN ADULTHOOD BACTERIAL MENINGITIS STUDY INVESTIGATORS*

ABSTRACT Background Mortality and morbidity rates are high among adults with acute bacterial meningitis, especially those with pneumococcal meningitis. In studies of bacterial meningitis in animals, adjuvant treatment



Dexamethasone - Adult Meningitis Study

Assessment of Outcome

The primary outcome measure was the score on the Glasgow Outcome Scale eight weeks after randomization, as assessed by the patient's physician. A score of 1 indicates death; 2, a vegetative state (the patient is unable to interact with the environment); 3, severe disability (the patient is unable to live independently but can follow commands); 4, moderate disability (the patient is capable of living independently but unable to return to work or school): and 5. mild

or no disability (the patient is able to return to work or school).¹² A favorable outcome was defined as a score of 5, and an unfavorable outcome as a score of 1 to 4. The Glasgow Outcome Scale has frequently been used in trials involving stroke and other brain injuries. It is a well-validated scale with good interobserver agreement.^{13,14}

N Engl J Med 2002;347:1549-56.

| Type of Data | Two Independent Samples | Related or Paired Samples | 3 or more Independent Samples | 3 or more Related Samples | Measures of Correlation |
|--------------|--|---|--|---------------------------------|--|
| Nominal | 1.Chi-square 2.Fisher's Exact | McNemar Test | Chi-square for k independe nt samples | Cochran Q | Contingency coefficient |
| Ordinal | 1.Mann- Whitney U 2.Wilcoxon Rank Sum | 1.Sign test 2.Wilcoxon Signed Rank | Kruskal- Wallis one way ANOVA | Freidman 2 way ANOVA | 1.Spearman 2.Kendal rank 3.Kendal Coe |
| Continuous | 1.Student's t-test 2.Mann- Whitney U | Paired t-test | 1-way ANOVA | 2-way ANOVA | Pearson's Correlation |



Statistical Analysis

 "Proportions of patients in the two groups were compared with Fisher's exact test. Two-tailed P values of less than 0.05 were considered to indicate statistical significance. Parametric and nonparametric values were tested with Student's t-test and the Mann–Whitney U test, respectively."

NEJM 2002;347(20):1549-56.

Main Results

| Outcome | Dexamethasone | Placebo | RR (95% CI) | P-value |
|-------------------|---------------|---------|-------------|---------|
| Unfavorable Outco | me | • | • | |
| All patients | 23/157 | 36/144 | | |
| S. pneumoniae | 15/58 | 26/50 | | |
| N. meningitidis | 4/5 | 5/47 | | |
| Other bacteria | 2/12 | 1/17 | | |
| Death | • | | • | |
| All patients | 11/157 | 21/144 | | |
| S. pneumoniae | 8/58 | 11/50 | | |
| N. meningitidis | 2/50 | 1/47 | | |
| Other bacteria | 1/12 | 1/17 | | |

Chi-squared vs. Fisher's exact

| Variable | Chi-square test | Fisher's exact test |
|------------------|--|--|
| Sample Size | Large | Small |
| Desired Accuracy | Approximate | "Exact" |
| Considerations | Becomes more accurate with larger sample sizes | More exact regardless of number but harder to calculate by hand using computer. Note: is it really "exact"? Typically used when > 20% of the cells have a frequency of < 5 because an approximation at this level is inadequate. |
| | | level is inadequate. |

MED REVIEWS



Main Results

| Outcome | Dexamethasone | Placebo | RR (95% CI) | P-value |
|-------------------|---------------|---------|-------------|---------|
| Unfavorable Outco | me | | | |
| All patients | 23/157 | 36/144 | | |
| S. pneumoniae | 15/58 | 26/50 | | |
| N. meningitidis | 4/5 | 5/47 | | |
| Other bacteria | 2/12 | 1/17 | | |
| Death | | | | |
| All patients | 11/157 | 21/144 | | |
| S. pneumoniae | 8/58 | 11/50 | | |
| N. meningitidis | 2/50 | 1/47 | | |
| Other bacteria | 1/12 | 1/17 | | |

NEJM 2002;347(20):1549-56.

HIGH-YIELD MED REVIEWS

Main Results

| Outcome | Dexamethasone | Placebo | RR (95% CI) | P-value | |
|---------------------|---------------|---------|-------------|---------|--|
| Unfavorable Outcome | | | | | |
| All patients | 23/157 | 36/144 | < 1 | | |
| S. pneumoniae | 15/58 | 26/50 | < 1 | | |
| N. meningitidis | 4/5 | 5/47 | < 1 | | |
| Other bacteria | 2/12 | 1/17 | > 1 | | |
| Death | • | | | | |
| All patients | 11/157 | 21/144 | < 1 | | |
| S. pneumoniae | 8/58 | 11/50 | < 1 | | |
| N. meningitidis | 2/50 | 1/47 | > 1 | | |
| Other bacteria | 1/12 | 1/17 | > 1 | | |

NEJM 2002;347(20):1549-56.

| HIGH-YIELD | | | | |
|------------|-------------|--|--|--|
| 2 | MED REVIEWS | | | |

Main Results

| Outcome | Dexamethasone | Placebo | RR (95% CI) | P-value |
|-------------------|---------------|---------|-------------|---------|
| Unfavorable Outco | me | • | • | |
| All patients | 23/157 | 36/144 | | |
| S. pneumoniae | 15/58 | 26/50 | | |
| N. meningitidis | 4/5 | 5/47 | | |
| Other bacteria | 2/12 | 1/17 | | |
| Death | | | • | • |
| All patients | 11/157 | 21/144 | | |
| S. pneumoniae | 8/58 | 11/50 | | |
| N. meningitidis | 2/50 | 1/47 | | |
| Other bacteria | 1/12 | 1/17 | | |

Relative Risk

- RR = incidence rate in exposed patients incidence rate in non-exposed patients
- RR = 1 (incidence is the same for both groups)
- RR = >1 (incidence in exposed group is higher)
- RR = <1 (incidence in exposed group is less)</p>

HIGH-YIELD MED REVIEWS

Main Results

RR = incidence rate in exposed patients incidence rate in non-exposed patients

1. Calculate the incidence in each group

HIGH-YIELD MED REVIEWS

Main Results

RR = incidence rate in exposed patients incidence rate in non-exposed patients

- 1. Calculate the incidence in each group
- 2. RR = ____ / ____ = 0.5





Main Results

| Outcome | Dexamethasone | Placebo | RR (95% CI) | P-value | |
|-------------------|---------------------|---------|--------------------|---------|--|
| Unfavorable Outco | Unfavorable Outcome | | | | |
| All patients | 23/157 | 36/144 | | | |
| S. pneumoniae | 15/58 | 26/50 | 0.50 (0.30 – 0.83) | | |
| N. meningitidis | 4/5 | 5/47 | | | |
| Other bacteria | 2/12 | 1/17 | | | |
| Death | | | | | |
| All patients | 11/157 | 21/144 | | | |
| S. pneumoniae | 8/58 | 11/50 | | | |
| N. meningitidis | 2/50 | 1/47 | | | |
| Other bacteria | 1/12 | 1/17 | | | |

Main Results

| Outcome | Dexamethasone | Placebo | RR (95% CI) | P-value |
|---------------------|---------------|---------|--------------------|---------|
| Unfavorable Outcome | | | | |
| All patients | 23/157 | 36/144 | 0.59 (0.37 – 0.94) | |
| S. pneumoniae | 15/58 | 26/50 | 0.50 (0.30 – 0.83) | |
| N. meningitidis | 4/5 | 5/47 | 0.75 (0.21 – 2.63) | |
| Other bacteria | 2/12 | 1/17 | 2.83 (0.29 – 27.8) | |
| Death | | | | |
| All patients | 11/157 | 21/144 | 0.48 (0.24 – 0.96) | |
| S. pneumoniae | 8/58 | 11/50 | 0.41 (0.19 – 0.86) | |
| N. meningitidis | 2/50 | 1/47 | 1.88 (0.76 – 20.1) | |
| Other bacteria | 1/12 | 1/17 | 1.42 (0.10 – 20.5) | |

Which results are significant?

NEJM 2002;347(20):1549-56.

HIGH-YIELD MED REVIEWS MED REVIEWS



NEJM 2002:347(20):1549-56.

How can we look at this another way?

NNT

RR = incidence rate in exposed patients incidence rate in non-exposed patients

- 1. Calculate the incidence in each group
- 2. RR = 0.26 / 0.52 = 0.5

=

- 3. ARR = ____ = ____
- 4. NNT = 1/
 - You would have to treat about _____ patients with dexamethasone 10 mg IV x 6 hrs x 4 days <u>with S. pneumonia</u> <u>meningitis</u> for 1 patient to have a favorable outcome.
 - Versus 10 patients if considering "all patients"

MED REVIEWS

The Results in the Context of Evidence

Cochrane Review

- 25 RCTs (n = 4121; with 2511 children, 1517 adults, 93 mixed):
- Quality of RCTs: 4 were high, 14 were medium; 7 were low — Results:
 - Non-significant reduction in mortality (regardless of age);
 17.8% vs. 19.9%, RR 0.9, 95% CI 0.8 1.01
 - Lower rates of severe hearing loss
 RR 0.67, 95% CI, 0.51 0.88
 - Lower rates of neurologic complications
 RR 0.83, 95% CI 0.69 1.00 **
 - Sub-group analysis only showed reduction in mortality if organism was Streptococcus pneumonia

 Not H. influenza or N. meningitidis

Agenda

- A General Overview
- Part 1 Conquering the Biostatistics Question
- Part 2 Interpreting Statistical Results Correctly
- A Special Coupon Code
- Live Q&A



Coupon

HIGH-YIELD MED REVIEWS

- Limited time coupon
 - -Coupon = BPSSAVE10
 - 10% OFF ENTIRE ORDER
 - Expires = Sept 30, 2022

Live Q&A



HIGH-YIELD MED REVIEWS

Why Should I Consider High-Yield Med Reviews? ...

What makes you different?



High Yield Study Tools



Knowledge Transfer

The High-Yield Approach

How does all of that fit together?





Knowledge Transfer



The High-Yield Approach



The High-Yield Approach



The High-Yield Approach

