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Module 6 Homework

1. **(60 points)** On the Golub et al. (1999) data, consider the “H4/j gene” gene (row 2972) and the “APS Prostate specific antigen” gene (row 2989). Setup the appropriate hypothesis for proving the following claims. Chose and carry out the appropriate tests.

- (a) The mean “H4/j gene” gene expression value in the ALL group is greater than -0.9 (note that this is negative 0.9).
- (b) The mean “H4/j gene” gene expression value in ALL group differs from the mean “H4/j gene” gene expression value in the AML group.
- (c) In the ALL group, the mean expression value for the “H4/j gene” gene is lower than the mean expression value for the “APS Prostate specific antigen” gene.
- (d) Let p_{H4j} denotes the proportion of patients for whom the “H4/j gene” expression values is greater than -0.6. We wish to show that p_{H4j} in the ALL group is less than 0.5.
- (e) The proportion p_{H4j} in the ALL group differs from the proportion p_{H4j} in the AML group.

You should state the hypothesis, show the R commands for the tests, show the output of these tests, and state your conclusion based on these outputs.

ANSWER:

CODE:

A

```
t_test_result <- t.test(data[data$group == "ALL", "H4_j_gene"], mu = -0.9,  
alternative = "greater")
```

Show the result

```
t_test_result
```

B

and a "H4_j_gene" column

```
all_group <- data[data$group == "ALL", "H4_j_gene"]
```



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```
aml_group <- data[data$group == "AML", "H4_j_gene"]

# Check the number of observations
n_all_group <- length(all_group)
n_aml_group <- length(aml_group)

cat("Number of observations in the ALL group:", n_all_group, "\n")
cat("Number of observations in the AML group:", n_aml_group, "\n")

# Check both groups have enough observations for the test
if (n_all_group >= 2 && n_aml_group >= 2) {
  # Conduct the t-test if both groups have enough data
  t_test_result <- t.test(all_group, aml_group)
  # Show the result
  t_test_result
} else {
  cat("One or both groups do not have enough data for the t-test.")
}

# C
# Create a data frame with a column
data <- data.frame(
  H4_j_gene = H4_j_gene_values,
  APS_Prostate_specific_antigen = APS_prostate_antigen_values,
  group = rep("ALL", length(H4_j_gene_values))
)

# Calculate a paired t-test
t_test_result <- t.test(data$H4_j_gene, data$APS_Prostate_specific_antigen,
  alternative = "less", paired = TRUE)

# Show the result
t_test_result
```



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#D

```
num_patients_H4j_ALL <- 10 # Replace with the actual number of patients in the  
"H4/j gene" group
```

```
num_patients_total_ALL <- 20 # Replace with the actual total number of patients  
in the "ALL" group
```

```
# Perform an exact binomial test for the one-sample proportion test
```

```
binom_test_result <- binom.test(num_patients_H4j_ALL,  
num_patients_total_ALL, p = 0.5, alternative = "less")
```

```
# Print the test result
```

```
print(binom_test_result)
```

#E

```
# Ensure that the counts are positive for both groups
```

```
num_patients_H4j_ALL <- 10 # Replace with the actual number of patients in the  
"H4/j gene" group
```

```
num_patients_total_ALL <- 20 # Replace with the actual total number of patients  
in the "ALL" group
```

```
# binomial test
```

```
binom_test_result <- binom.test(num_patients_H4j_ALL,  
num_patients_total_ALL, p = 0.5, alternative = "less")
```

```
# Print the test result
```

```
print(binom_test_result)
```

2. (10 points) Suppose that the probability to reject a biological hypothesis by the results of a certain experiment is 0.03. This experiment is repeated 3000 times.

ANSWER:

CODE:



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a) Calculate the expected rejections

```
probability_rejection = 0.03
```

```
num_simulations = 3000
```

```
expected_rejections = probability_rejection * num_simulations
```

```
cat("Expected Rejections:", expected_rejections, "\n")
```

b) Calculate the probability of less than 75 rejections

```
num_rejections = 74
```

```
probability_less_than_75_rejections = pbinom(num_rejections, size =  
num_simulations, prob = probability_rejection)
```

```
cat("Probability of Less Than 75 Rejections:", probability_less_than_75_rejections,  
"\n")
```

(a) How many rejections do you expect?

CODE:

a) Calculate the expected rejections

```
probability_rejection = 0.03
```

```
num_simulations = 3000
```

```
expected_rejections = probability_rejection * num_simulations
```

```
cat("Expected Rejections:", expected_rejections, "\n")
```

OUTPUT: Expected Rejections: 90

(b) What is the probability of less than 75 rejections?

CODE:

#b) Calculate the probability of less than 75 rejections

```
num_rejections = 74
```

```
probability_less_than_75_rejections = pbinom(num_rejections, size =  
num_simulations, prob = probability_rejection)
```

```
cat("Probability of Less Than 75 Rejections:", probability_less_than_75_rejections,  
"\n")
```



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OUTPUT: Probability of Less Than 75 Rejections: 0.04537989

3. (10 points)

For testing $H_0: \mu=5$ versus $H_A: \mu>5$, we consider a new $\alpha=0.1$ level test which rejects when $t_{obs} = \frac{\bar{X}-5}{s/\sqrt{n}}$ falls between $t_{0.3,n-1}$ and $t_{0.4,n-1}$.

Use a **Monte Carlo simulation** to estimate the Type I error rate of this test when $n=30$. Do 10,000 simulation runs of data sets from the $N(\mu = 5, \sigma = 4)$. Please show the R script for the simulation, and the R outputs for running the script. Provide your numerical estimate for the Type I error rate. Is this test valid (that is, is its Type I error rate same as the nominal $\alpha=0.1$ level)?

ANSWER:

CODE:

```
# Parameters
```

```
alpha <- 0.1
```

```
datasets_simulations <- 10000
```

```
n <- 30
```

```
null_mean <- 5
```

```
# Function to perform a single simulation and return 1 if Type I error occurs, 0 otherwise
```

```
simulation <- function() {
```

```
  data_sample <- rnorm(n, mean = null_mean)
```

```
  t_test_result <- t.test(data_sample, mu = 5, alternative = "greater")
```

```
  return(as.numeric(t_test_result$p.value < alpha))
```

```
}
```



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```
# Run simulations and count Type I errors
errors <- sum(replicate(datasets_simulations, simulation()))

# Calculate the estimated Type I error rate
type_I_error_rate <- errors / datasets_simulations

cat("Estimated Type I Error Rate:", type_I_error_rate)
```

OUTPUT: Estimated Type I Error Rate: 0.0979

4. (20 points)

On the Golub et al. (1999) data set, do **Welch two-sample t-tests** to compare every gene's expression values in ALL group versus in AML group.

- (a) Use Bonferroni and FDR adjustments both at 0.05 level. How many genes are differentially expressed according to these two criteria?
- (b) Find the gene names for the top three strongest differentially expressed genes (i.e., minimum p-values). Hint: the gene names are stored in ***golub.gnames***.

Please submit your R commands together with your answers to each part of the question.

ANSWER:

CODE:

```
# Verify that the gene expression data has been appropriately put into the
"golub_data" data frame.
# Verify that your gene names are in 'golub.gnames'.
```



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```
t_test_results <- lapply(names(golub_data), function(gene_name) {  
  gene_expression <- golub_data[[gene_name]]  
  group_all <- gene_expression[golub_data$group == 'ALL']  
  group_aml <- gene_expression[golub_data$group == 'AML']  
  t_test_result <- t.test(group_all, group_aml)  
  t_test_result$p.value  
})
```

```
alpha <- 0.05
```

```
bonferroni_cutoff <- alpha / length(t_test_results)  
bonferroni_adjusted <- p.adjust(t_test_results, method = "bonferroni")
```

```
fdr_adjusted <- p.adjust(t_test_results, method = "fdr")
```

```
differentially_expressed_genes_bonferroni <- sum(bonferroni_adjusted < alpha)  
differentially_expressed_fdr <- sum(fdr_adjusted < alpha)
```

```
cat("Differentially Expressed Genes (Bonferroni):",  
differentially_expressed_genes_bonferroni, "\n")  
cat("Differentially Expressed Genes (FDR):", differentially_expressed_fdr, "\n")
```

```
top_three_genes <- order(t_test_results)[1:3]  
top_gene_names <- names(golub_data)[top_three_genes]
```

```
cat("Top Three Differentially Expressed Genes:", top_gene_names, "\n")
```