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Automating Reviews in Medicine

automating reviews in medicine

The medical literature is enormous. Pubmed, a database of medical publications maintained by the U.S. National Library of Medicine, has indexed over 23 million medical publications. Further, the rate of medical publication has increased over time, and now there are nearly 1 million new publications in the field each year, or more than one per minute.

The large size and fast-changing nature of the medical literature has increased the need for reviews, which search databases like Pubmed for papers on a particular topic and then report results from the papers found. While such reviews are often performed manually, with multiple people reviewing each search result, this is tedious and time consuming. In this problem, we will see how text analytics can be used to automate the process of information retrieval.

The dataset consists of the titles (variable *title*) and abstracts (variable *abstract*) of papers retrieved in a [Pubmed](#) search. Each search result is labeled with whether the paper is a clinical trial testing a drug therapy for cancer (variable *trial*). These labels were obtained by two people reviewing each search result and accessing the actual paper if necessary, as part of a literature review of clinical trials testing drug therapies for advanced and metastatic breast cancer.

Problem 1.1 - Loading the Data

1 point possible (graded)

Load [clinical_trial.csv](#) into a data frame called trials (remembering to add the

argument `stringsAsFactors=FALSE`), and investigate the data frame with `summary()` and `str()`.

IMPORTANT NOTE: Some students have been getting errors like "invalid multibyte string" when performing certain parts of this homework question. If this is happening to you, use the argument `fileEncoding="latin1"` when reading in the file with `read.csv`. This should cause those errors to go away.

We can use R's string functions to learn more about the titles and abstracts of the located papers. The `nchar()` function counts the number of characters in a piece of text. Using the `nchar()` function on the variables in the data frame, answer the following questions:

How many characters are there in the longest abstract? (Longest here is defined as the abstract with the largest number of characters.)

Answer: 3708

Explanation

You can load the data set into R with the following command:

```
trials = read.csv("clinical_trial.csv", stringsAsFactors=FALSE)
```

From `summary(nchar(trials$abstract))` or `max(nchar(trials$abstract))`, we can read the maximum length.

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You have used 0 of 3 attempts

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Problem 1.2 - Loading the Data

1 point possible (graded)

How many search results provided no abstract? (HINT: A search result provided no abstract if the number of characters in the abstract field is zero.)

Answer: 112

Explanation

From `table(nchar(trials$abstract) == 0)` or `sum(nchar(trials$abstract) == 0)`, we can find the number of missing abstracts.

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Problem 1.3 - Loading the Data

1 point possible (graded)

Find the observation with the minimum number of characters in the title (the variable "title") out of all of the observations in this dataset. What is the text of the title of this article? Include capitalization and punctuation in your response, but don't include the quotes.

Answer: A decade of letrozole: FACE.

Explanation

To identify which title is the shortest, we can use `which.min(nchar(trials$title))`

From this, we know the 1258th title is the shortest. We can access this title with `trials$title[1258]`.

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Problem 2.1 - Preparing the Corpus

0.0/4.0 points (graded)

Because we have both title and abstract information for trials, we need to build two corpora instead of one. Name them `corpusTitle` and `corpusAbstract`.

Following the commands from lecture, perform the following tasks (you might need to load the "tm" package first if it isn't already loaded). Make sure to perform them in this order.

1) Convert the title variable to `corpusTitle` and the abstract variable to `corpusAbstract`.

2) Convert `corpusTitle` and `corpusAbstract` to lowercase.

- 3) Remove the punctuation in corpusTitle and corpusAbstract.
- 4) Remove the English language stop words from corpusTitle and corpusAbstract.
- 5) Stem the words in corpusTitle and corpusAbstract (each stemming might take a few minutes).
- 6) Build a document term matrix called dtmTitle from corpusTitle and dtmAbstract from corpusAbstract.
- 7) Limit dtmTitle and dtmAbstract to terms with sparseness of at most 95% (aka terms that appear in at least 5% of documents).
- 8) Convert dtmTitle and dtmAbstract to data frames (keep the names dtmTitle and dtmAbstract).

If the code `length(stopwords("english"))` does not return 174 for you, then please run the line of code [in this file](#), which will store the standard stop words in a variable called `sw`. When removing stop words, use `tm_map(corpusTitle, removeWords, sw)` and `tm_map(corpusAbstract, removeWords, sw)` instead of `tm_map(corpusTitle, removeWords, stopwords("english"))` and `tm_map(corpusAbstract, removeWords, stopwords("english"))`.

Explanation

Below we provide the code for corpusTitle; only minor modifications are needed to build corpusAbstract.

```
corpusTitle = VCorpus(VectorSource(trials$title))
corpusTitle = tm_map(corpusTitle, content_transformer(tolower))
corpusTitle = tm_map(corpusTitle, removePunctuation)
corpusTitle = tm_map(corpusTitle, removeWords, stopwords("english"))
corpusTitle = tm_map(corpusTitle, stemDocument)
dtmTitle = DocumentTermMatrix(corpusTitle)
dtmTitle = removeSparseTerms(dtmTitle, 0.95)
dtmTitle = as.data.frame(as.matrix(dtmTitle))
```

How many terms remain in dtmTitle after removing sparse terms (aka how many columns does it have)?

Answer: 31

How many terms remain in dtmAbstract?

Answer: 335

Explanation

These can be read from `str(dtmTitle)` and `str(dtmAbstract)`. Other than `str()`, the `dim()` or `ncol()` functions could have been used. If you used `fileEncoding="latin1"` when reading in the datafile, you'll have a few extra terms in `dtmAbstract`, but you should get the answer correct.

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Problem 2.2 - Preparing the Corpus

1 point possible (graded)

What is the most likely reason why `dtmAbstract` has so many more terms than `dtmTitle`?

- ☒ Abstracts tend to have many more words than titles ✓
- ☐ Abstracts tend to have a much wider vocabulary than titles
- ☐ More papers have abstracts than titles

Explanation

Because titles are so short, a word needs to be very common to appear in 5% of titles. Because abstracts have many more words, a word can be much less common and still appear in 5% of abstracts.

While abstracts may have wider vocabulary, this is a secondary effect. As we saw in the previous subsection, all papers have titles, but not all have abstracts.

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Problem 2.3 - Preparing the Corpus

1 point possible (graded)

What is the most frequent word stem across all the abstracts? Hint: you can use `colSums()` to compute the frequency of a word across all the abstracts.

Answer: patient

Explanation

We can compute the column sums and then identify the most common one with:

```
csAbstract = colSums(dtmAbstract)
```

```
which.max(csAbstract)
```

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Problem 3.1 - Building a model

1 point possible (graded)

We want to combine `dtmTitle` and `dtmAbstract` into a single data frame to make predictions. However, some of the variables in these data frames have the same names. To fix this issue, run the following commands:

```
colnames(dtmTitle) = paste0("T", colnames(dtmTitle))
```

```
colnames(dtmAbstract) = paste0("A", colnames(dtmAbstract))
```

What was the effect of these functions?

- ☐ Removing the words that are in common between the titles and the abstracts.
- ☒ Adding the letter T in front of all the title variable names and adding the letter A in front of all the abstract variable names. ✓
- ☐ Adding the letter T in front of all the title variable names that also appear in the abstract data frame, and adding an A in front of all the abstract variable names that appear in the title data frame.

Explanation

The first line pastes a T at the beginning of each column name for dtmTitle, which are the variable names. The second line does something similar for the Abstract variables - it pastes an A at the beginning of each column name for dtmAbstract, which are the variable names.

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Problem 3.2 - Building a Model

1 point possible (graded)

Using cbind(), combine dtmTitle and dtmAbstract into a single data frame called dtm:

```
dtm = cbind(dtmTitle, dtmAbstract)
```

As we did in class, add the dependent variable "trial" to dtm, copying it from the original data frame called trials. How many columns are in this combined data frame?

Answer: 367

Explanation

The combination can be accomplished with:

```
dtm = cbind(dtmTitle, dtmAbstract)
```

```
dtm$trial = trials$trial
```

The number of variables in the combined data frame can be read from `str(dtm)` or `ncol(dtm)`. If you used `fileEncoding="latin1"` when reading in the file, you should have 5 extra variables (but the answer should be graded as correct).

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Problem 3.3 - Building a Model

1 point possible (graded)

Now that we have prepared our data frame, it's time to split it into a training and testing set and to build regression models. Set the random seed to 144 and use the `sample.split` function from the `caTools` package to split `dtm` into data frames named "train" and "test", putting 70% of the data in the training set.

Explanation

This can be accomplished with:

```
set.seed(144)
```

```
spl = sample.split(dtm$trial, 0.7)
```

```
train = subset(dtm, spl == TRUE)
```

```
test = subset(dtm, spl == FALSE)
```

What is the accuracy of the baseline model on the training set? (Remember that the baseline model predicts the most frequent outcome in the training set for all observations.)

Answer: 0.561

Explanation

Just as in any binary classification problem, the naive baseline always predicts the most common class. From `table(train$trial)`, we see 730 training set results were not trials, and 572 were trials. Therefore, the naive baseline always predicts a result is not a trial, yielding accuracy of $730/(730+572)$.

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You have used 0 of 4 attempts

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Problem 3.4 - Building a Model

0.0/2.0 points (graded)

Build a CART model called `trialCART`, using all the independent variables in the training set to train the model, and then plot the CART model. Just use the default parameters to build the model (don't add a `minbucket` or `cp` value). Remember to add the `method="class"` argument, since this is a classification problem.

What is the name of the first variable the model split on?

Answer: Tphase

Explanation

This can be accomplished with:

```
trialCART = rpart(trial~., data=train, method="class")  
prp(trialCART)
```

The first split checks whether or not Tphase is less than 0.5

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Problem 3.5 - Building a Model

1 point possible (graded)

Obtain the training set predictions for the model (do not yet predict on the test set). Extract the predicted probability of a result being a trial (recall that this involves not setting a `type` argument, and keeping only the second column of the `predict` output). What is the maximum predicted probability for any result?

Answer: 0.872

Explanation

The training set predictions can be obtained and summarized with the following commands:

```
predTrain = predict(trialCart)[,2]
summary(predTrain)
```

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Problem 3.6 - Building a Model

1 point possible (graded)

Without running the analysis, how do you expect the maximum predicted probability to differ in the testing set?

- ☐ The maximum predicted probability will likely be smaller in the testing set.
- ☒ The maximum predicted probability will likely be exactly the same in the testing set. ✓
- ☐ The maximum predicted probability will likely be larger in the testing set.

Explanation

Because the CART tree assigns the same predicted probability to each leaf node and there are a small number of leaf nodes compared to data points, we expect exactly the same maximum predicted probability.

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Problem 3.7 - Building a Model

3 points possible (graded)

For these questions, use a threshold probability of 0.5 to predict that an observation is a clinical trial.

What is the training set accuracy of the CART model?

Answer: 0.823

What is the training set sensitivity of the CART model?

Answer: 0.771

What is the training set specificity of the CART model?

Answer: 0.864

Explanation

We can compare the predictions with threshold 0.5 to the true results in the training set with:

```
table(train$trial, predTrain >= 0.5)
```

From this, we read the following confusion matrix (rows are true outcome, columns are predicted outcomes):

	FALSE	TRUE
0	631	99
1	131	441

We conclude that the model has training set accuracy

$(631+441)/(631+441+99+131)$, sensitivity $441/(441+131)$ and specificity $631/(631+99)$.

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Problem 4.1 - Evaluating the model on the testing set

0.0/2.0 points (graded)

Evaluate the CART model on the testing set using the predict function and creating a vector of predicted probabilities predTest.

What is the testing set accuracy, assuming a probability threshold of 0.5 for predicting that a result is a clinical trial?

Answer: 0.758

Explanation

The testing set predictions can be obtained and compared to the true outcomes with:

```
predTest = predict(trialCART, newdata=test)[,2]
```

```
table(test$trial, predTest >= 0.5)
```

This yields the following confusion matrix:

```
FALSE TRUE
```

```
0 261 52
```

```
1 83 162
```

From this, we read that the testing set accuracy is $(261+162)/(261+162+83+52)$.

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Problem 4.2 - Evaluating the Model on the Testing Set

0.0/2.0 points (graded)

Using the ROCR package, what is the testing set AUC of the prediction model?

Answer: 0.837

Explanation

The AUC can be determined using the following code:

```
library(ROCR)
```

```
pred = prediction(predTest, test$trial)
```

```
as.numeric(performance(pred, "auc")@y.values)
```

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part 5: decision-maker tradeoffs

The decision maker for this problem, a researcher performing a review of the

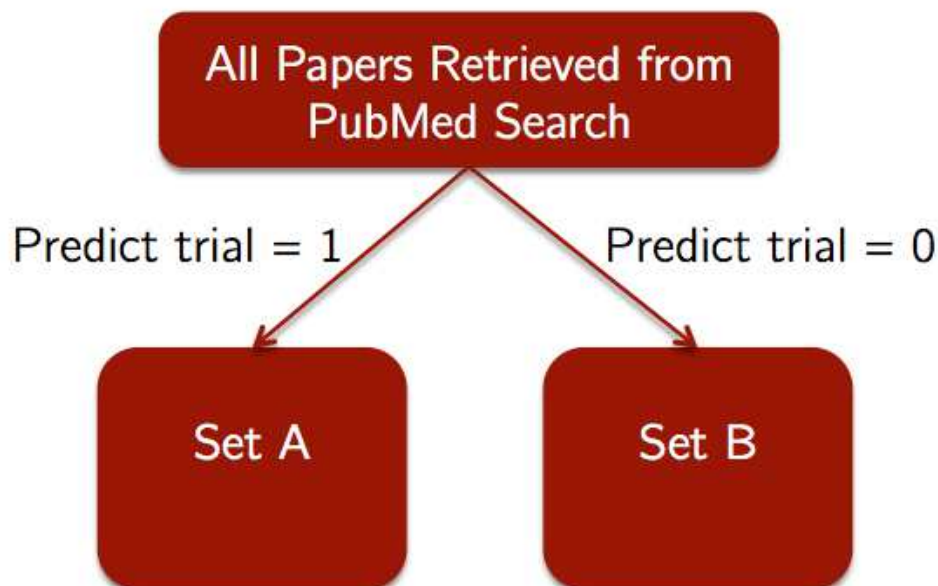
medical literature, would use a model (like the CART one we built here) in the following workflow:

1) For all of the papers retrieved in the PubMed Search, predict which papers are clinical trials using the model. This yields some initial Set A of papers predicted to be trials, and some Set B of papers predicted not to be trials. (See the figure below.)

2) Then, the decision maker manually reviews all papers in Set A, verifying that each paper meets the study's detailed inclusion criteria (for the purposes of this analysis, we assume this manual review is 100% accurate at identifying whether a paper in Set A is relevant to the study). This yields a more limited set of papers to be included in the study, which would ideally be all papers in the medical literature meeting the detailed inclusion criteria for the study.

3) Perform the study-specific analysis, using data extracted from the limited set of papers identified in step 2.

This process is shown in the figure below.



Problem 5.1 - Decision-Maker Tradeoffs

1 point possible (graded)

What is the cost associated with the model in Step 1 making a false negative prediction?

- ☐ A paper will be mistakenly added to Set A, yielding additional work in Step 2 of the process but not affecting the quality of the results of Step 3.
- ☐ A paper will be mistakenly added to Set A, definitely affecting the quality of the results of Step 3.
- ☒ A paper that should have been included in Set A will be missed, affecting the quality of the results of Step 3. ✓
- ☐ There is no cost associated with a false negative prediction.

Explanation

By definition, a false negative is a paper that should have been included in Set A but was missed by the model. This means a study that should have been included in Step 3 was missed, affecting the results.

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Problem 5.2 - Decision-Maker Tradeoffs

1 point possible (graded)

What is the cost associated with the model in Step 1 making a false positive prediction?

- ☒ A paper will be mistakenly added to Set A, yielding additional work in Step 2 of the process but not affecting the quality of the results of Step 3. ✓
- ☐ A paper will be mistakenly added to Set A, definitely affecting the quality of the results of Step 3.
- ☐ A paper that should have been included in Set A will be missed, affecting the quality of the results of Step 3.
- ☐ There is no cost associated with a false positive prediction.

Explanation

By definition, a false positive is a paper that should not have been included in Set A but that was actually included. However, because the manual review in Step 2 is assumed to be 100% effective, this extra paper will not make it into the more limited set of papers, and therefore this mistake will not affect the analysis in Step 3.

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Problem 5.3 - Decision-Maker Tradeoffs

1 point possible (graded)

Given the costs associated with false positives and false negatives, which of the following is most accurate?

- ☐ A false positive is more costly than a false negative; the decision maker should use a probability threshold greater than 0.5 for the machine learning model.
- ☐ A false positive is more costly than a false negative; the decision maker should use a probability threshold less than 0.5 for the machine learning model.
- ☐ A false negative is more costly than a false positive; the decision maker should use a probability threshold greater than 0.5 for the machine learning model.
- ☒ A false negative is more costly than a false positive; the decision maker should use a probability threshold less than 0.5 for the machine learning model. ✓

Explanation

A false negative might negatively affect the results of the literature review and analysis, while a false positive is a nuisance (one additional paper that needs to be manually checked). As a result, the cost of a false negative is much higher than the cost of a false positive, so much so that many studies actually use no machine learning (aka no Step 1) and have two people manually review each search result in Step 2. As always, we prefer a lower threshold in cases where false negatives are more costly than false positives, since we will make fewer negative predictions.

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