#### Class 18 Pertussis Mini Project

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#### Is Pertussis on the rise

Install packages - datapasta

don't need to Load library! We can click Addins button instead and paste as data frame - one way to scrape data from the internet. Downside is, we need to make sure to copy/paste accurately

Data copied from: https://www.cdc.gov/pertussis/surv-reporting/cases-by-year.html

```
1991L, 1992L, 1993L, 1994L, 1995L, 1996L,
                                      1997L, 1998L, 1999L, 2000L, 2001L,
                                      2002L,2003L,2004L,2005L,2006L,2007L,
                                      2008L,2009L,2010L,2011L,2012L,
                                      2013L, 2014L, 2015L, 2016L, 2017L, 2018L,
                                      2019L),
  Cases = c(107473, 164191, 165418, 152003,
                                      202210, 181411, 161799, 197371,
                                      166914, 172559, 215343, 179135, 265269,
                                      180518, 147237, 214652, 227319, 103188,
                                      183866,222202,191383,191890,109873,
                                      133792,109860,156517,74715,69479,
                                      120718,68687,45030,37129,60886,
                                     62786,31732,28295,32148,40005,
                                      14809, 11468, 17749, 17135, 13005, 6799,
                                     7717,9718,4810,3285,4249,3036,
                                     3287,1759,2402,1738,1010,2177,2063,
                                      1623,1730,1248,1895,2463,2276,
                                     3589,4195,2823,3450,4157,4570,
                                      2719,4083,6586,4617,5137,7796,6564,
                                     7405,7298,7867,7580,9771,11647,
                                     25827, 25616, 15632, 10454, 13278,
                                      16858, 27550, 18719, 48277, 28639, 32971,
                                      20762,17972,18975,15609,18617)
)
```

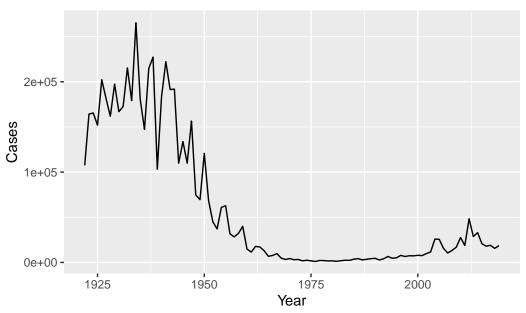
Q1. With the help of the R "addin" package datapasta assign the CDC pertussis case number data to a data frame called cdc and use ggplot to make a plot of cases numbers over time.

```
library(ggplot2)

baseplot <- ggplot(data = cdc) +
   aes(x = Year, y = Cases) +
   geom_line() +
   labs(title = "Pertussis Cases Over Time in the US")

baseplot</pre>
```

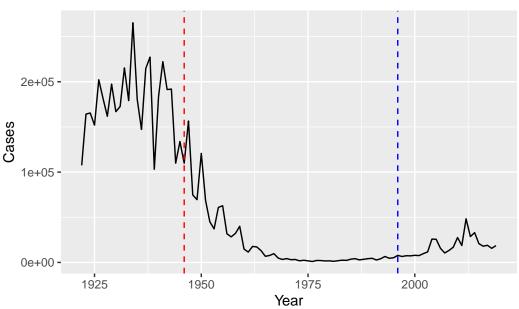
#### Pertussis Cases Over Time in the US



# Q2. Using the ggplot geom\_vline() function add lines to your previous plot for the 1946 introduction of the wP vaccine and the 1996 switch to aP vaccine (see example in the hint below). What do you notice?

```
#Adding a vline for the introduction of the first pertussis vaccine in 1996 and the change
baseplot +
  geom_vline(xintercept = 1946, color = "red", linetype = 2) + geom_vline(xintercept = 1996)
```

#### Pertussis Cases Over Time in the US



## Q3. Describe what happened after the introduction of the aP vaccine? Do you have a possible explanation for the observed trend?

The pertussis strain could be mutating and the vaccine formulation could be less effective; alternately, the new vaccine formulation could be less effective versus the one formulated from the whole bacteria. Or, it could be a decrease in vaccination rates due to the paper from 2006. Evidence suggests that it's not variation in the bacteria that causes pertussis, but it might be waning efficacy due to the new formulation of the pertussis vaccine.

Installing the jsonlite package Loading in the library

```
library(jsonlite)
```

1

Reading a json file from the website:

```
subject <- read_json("https://www.cmi-pb.org/api/subject", simplifyVector = TRUE)
head(subject, 3)

subject_id infancy_vac biological_sex ethnicity race</pre>
```

Female Not Hispanic or Latino White

```
2
           2
                                  Female Not Hispanic or Latino White
                      wP
3
           3
                      wP
                                 Female
                                                        Unknown White
 year_of_birth date_of_boost
                                    dataset
     1986-01-01
                   2016-09-12 2020_dataset
2
     1968-01-01
                   2019-01-28 2020_dataset
     1983-01-01
                   2016-10-10 2020_dataset
```

### Q4. How many aP and wP infancy vaccinated subjects are in the dataset?

```
wP: 49, aP = 47
table(subject$infancy_vac)
aP wP
47 49
```

### Q5. How many Male and Female subjects/patients are in the dataset?

```
F: 66 M: 30

table(subject$biological_sex)

Female Male
66 30
```

### Q6. What is the breakdown of race and biological sex (e.g. number of Asian females, White males etc...)?

```
table(subject$race, subject$biological_sex)
```

|   | Female | Male |
|---|--------|------|
| American Indian/Alaska Native             | 0      | 1    |
| Asian                                     | 18     | 9    |
| Black or African American                 | 2      | 0    |
| More Than One Race                        | 8      | 2    |
| Native Hawaiian or Other Pacific Islander | 1      | 1    |
| Unknown or Not Reported                   | 10     | 4    |
| White                                     | 27     | 13   |

#### **Dates**

```
head(subject$year_of_birth, 3)

[1] "1986-01-01" "1968-01-01" "1983-01-01"

head(subject$date_of_boost, 3)

[1] "2016-09-12" "2019-01-28" "2016-10-10"

library(lubridate)

Loading required package: timechange

Attaching package: 'lubridate'

The following objects are masked from 'package:base':
   date, intersect, setdiff, union

today()

[1] "2022-11-30"

today() - ymd("2000-01-01")
```

Time difference of 8369 days

```
time_length( today() - ymd("2000-01-01"), "years")

[1] 22.91307

Making a new column for age of all subjects in the trial

subject$age <- time_length( today() - ymd(subject$year_of_birth), "years")
head(subject, 3)

subject_id infancy_vac biological_sex ethnicity race</pre>
```

```
ethnicity race
1
           1
                     wΡ
                                Female Not Hispanic or Latino White
2
           2
                     wP
                                Female Not Hispanic or Latino White
3
           3
                     wP
                                Female
                                                       Unknown White
 year_of_birth date_of_boost
                                   dataset
     1986-01-01
                  2016-09-12 2020_dataset 36.91170
1
     1968-01-01
                  2019-01-28 2020_dataset 54.91307
2
     1983-01-01
                  2016-10-10 2020_dataset 39.91239
3
```

## Q7. Using this approach determine (i) the average age of wP individuals, (ii) the average age of aP individuals; and (iii) are they significantly different?

```
summary(subject$age)

Min. 1st Qu. Median Mean 3rd Qu. Max.
22.91 25.91 27.91 30.77 34.91 54.91
```

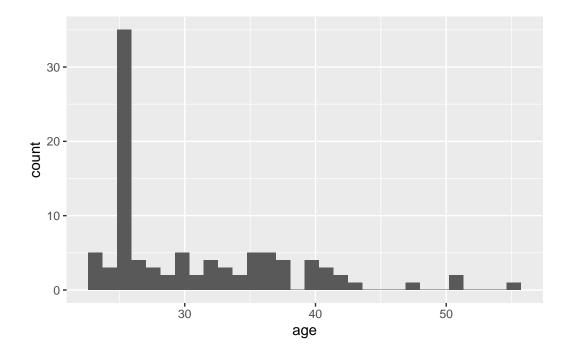
Q8:

Q9. With the help of a faceted boxplot or histogram (see below), do you think these two groups are significantly different?

Making a histogram

```
base_hist <- ggplot(subject) +
  aes(age) +
  geom_histogram()
base_hist</pre>
```

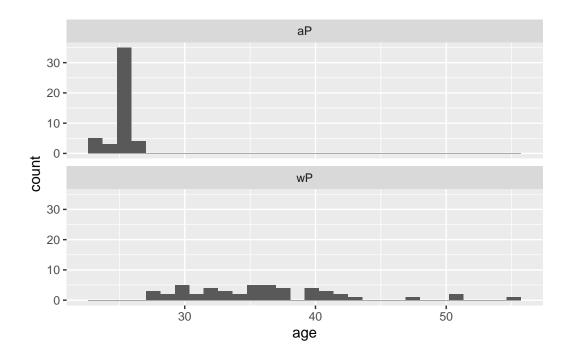
`stat\_bin()` using `bins = 30`. Pick better value with `binwidth`.



Facet wrapping by type of vaccine shows the point at which the aP vaccine was introduced

```
base_hist +
  facet_wrap(vars(infancy_vac), nrow = 2)
```

`stat\_bin()` using `bins = 30`. Pick better value with `binwidth`.



Using dplyr to join different tables together - get specimen table and join with subject to have all information in one place

```
library(dplyr)
```

```
Attaching package: 'dplyr'
```

The following objects are masked from 'package:stats':

```
filter, lag
```

The following objects are masked from 'package:base':

```
intersect, setdiff, setequal, union
```

Read the specimen table

```
specimen <- read_json("https://www.cmi-pb.org/api/specimen", simplifyVector = TRUE)</pre>
```

look at specimen table:

```
head(specimen, 3)
```

```
specimen_id subject_id actual_day_relative_to_boost
2
            2
                                                    736
                        1
3
                        1
 planned_day_relative_to_boost specimen_type visit
                                          Blood
1
                               0
2
                             736
                                          Blood
                                                   10
3
                               1
                                          Blood
                                                    2
```

#### Q9. Complete the code to join specimen and subject tables to make a new merged data frame containing all specimen records along with their associated subject details:

1

Using a join function to bring the tables together. It will automatically find what's in common between the tables, which here is "subject\_id", and joins them together - same number of rows as in specimens, but with extra columns from subject

```
library(dplyr)
  meta <- inner_join(specimen, subject)</pre>
Joining, by = "subject_id"
  dim(meta)
[1] 729 14
Now reading some experimental data
  titer <- read_json("https://www.cmi-pb.org/api/ab_titer", simplifyVector = TRUE)</pre>
  head(titer)
```

```
specimen_id isotype is_antigen_specific antigen
                                                          MFI MFI_normalised
1
            1
                                     FALSE
                                             Total 1110.21154
                                                                     2.493425
            1
2
                                     FALSE
                                             Total 2708.91616
                                                                     2.493425
                  IgE
3
            1
                                      TRUE
                                                PΤ
                  IgG
                                                     68.56614
                                                                     3.736992
4
            1
                  IgG
                                      TRUE
                                               PRN 332.12718
                                                                     2.602350
5
            1
                  IgG
                                      TRUE
                                               FHA 1887.12263
                                                                    34.050956
                  IgE
                                      TRUE
                                               ACT
                                                      0.10000
                                                                    1.000000
  unit lower_limit_of_detection
1 UG/ML
                        2.096133
2 IU/ML
                       29.170000
3 IU/ML
                        0.530000
4 IU/ML
                        6.205949
5 IU/ML
                        4.679535
6 IU/ML
                        2.816431
```

## Q10. Now using the same procedure join meta with titer data so we can further analyze this data in terms of time of visit aP/wP, male/female etc.

```
Join the titer table with the meta table!
```

```
abdata <- inner_join(meta, titer)

Joining, by = "specimen_id"

dim(abdata)

[1] 32675 21</pre>
```

### Q11. How many specimens (i.e. entries in abdata) do we have for each isotype?

```
table(abdata$isotype)

IgE IgG IgG1 IgG2 IgG3 IgG4
6698 1413 6141 6141 6141 6141
```

### Q12. What do you notice about the number of visit 8 specimens compared to other visits?

Much fewer of them! So, we will drop this from our data, as well as focus in on one isotype - I will use IgG1

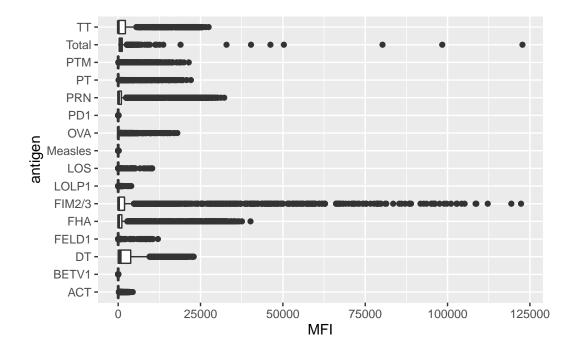
```
table(abdata$visit)

1 2 3 4 5 6 7 8
5795 4640 4640 4640 4640 4320 3920 80
```

Making a plot for titer levels of different antiboides

```
ggplot(abdata) +
  aes(MFI, antigen) +
  geom_boxplot()
```

Warning: Removed 1 rows containing non-finite values (stat\_boxplot).

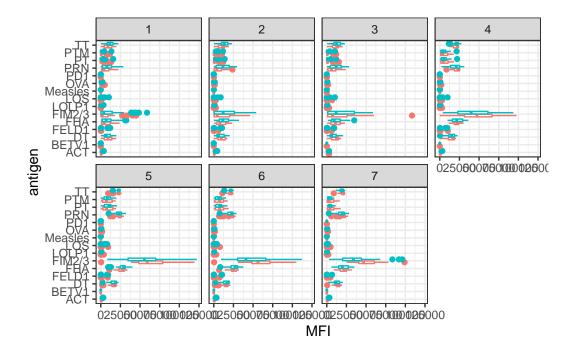


Will define ig1 as dataset for only IgG1 and without visit 8:

```
ig1 <- abdata %>%
    filter(isotype == "IgG1", visit != 8)
  head(ig1)
  specimen_id subject_id actual_day_relative_to_boost
1
                        1
                                                      -3
2
            1
                        1
                                                      -3
3
            1
                        1
                                                      -3
4
            1
                        1
                                                      -3
5
                                                      -3
            1
                        1
                                                      -3
  planned_day_relative_to_boost specimen_type visit infancy_vac biological_sex
                               0
                                          Blood
                                                    1
                                                                wP
                                                                            Female
1
2
                               0
                                          Blood
                                                    1
                                                                wP
                                                                            Female
3
                               0
                                                                            Female
                                          Blood
                                                    1
                                                                wP
4
                               0
                                                     1
                                                                            Female
                                          Blood
                                                                wP
5
                               0
                                          Blood
                                                    1
                                                                wP
                                                                            Female
6
                               0
                                                                wΡ
                                                                            Female
                                          Blood
               ethnicity race year_of_birth date_of_boost
                                                                   dataset
                                                                                age
1 Not Hispanic or Latino White
                                    1986-01-01
                                                  2016-09-12 2020_dataset 36.9117
2 Not Hispanic or Latino White
                                    1986-01-01
                                                  2016-09-12 2020_dataset 36.9117
3 Not Hispanic or Latino White
                                                  2016-09-12 2020_dataset 36.9117
                                    1986-01-01
4 Not Hispanic or Latino White
                                                  2016-09-12 2020_dataset 36.9117
                                    1986-01-01
5 Not Hispanic or Latino White
                                    1986-01-01
                                                  2016-09-12 2020 dataset 36.9117
6 Not Hispanic or Latino White
                                    1986-01-01
                                                  2016-09-12 2020_dataset 36.9117
  isotype is_antigen_specific antigen
                                               MFI MFI normalised unit
1
     IgG1
                          TRUE
                                    ACT 274.355068
                                                         0.6928058 IU/ML
2
     IgG1
                          TRUE
                                   LOS 10.974026
                                                         2.1645083 IU/ML
3
     IgG1
                          TRUE
                                 FELD1
                                          1.448796
                                                         0.8080941 IU/ML
4
                                 BETV1
     IgG1
                          TRUE
                                          0.100000
                                                         1.0000000 IU/ML
5
                                          0.100000
     IgG1
                                 LOLP1
                                                         1.0000000 IU/ML
                          TRUE
     IgG1
                          TRUE Measles
                                         36.277417
                                                         1.6638332 IU/ML
  lower_limit_of_detection
1
                   3.848750
2
                   4.357917
3
                   2.699944
4
                   1.734784
5
                   2.550606
6
                   4.438966
```

### Q13. Complete the following code to make a summary boxplot of Ab titer levels for all antigens:

```
ggplot(ig1) +
  aes(MFI, antigen, col = infancy_vac) +
  geom_boxplot(show.legend = FALSE) +
  facet_wrap(vars(visit), nrow=2) +
  theme_bw()
```



### Q14. What antigens show differences in the level of IgG1 antibody titers recognizing them over time? Why these and not others?

TT, PTM, PT, PRN, FIM2/3, LOS, FHA, FELD1 and DT

These are all antigens in the vaccine, whereas antibodies to measles don't change because people in the study haven't been exposed to measles