Class15 Pertussis and the CMI-PB project

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key words: web scrapping, cmi, json, dplyr, inner_join(), filter(), facet_wrap(),

Pertussis a.k.a Whooping cough, is a highly contagious lung infection caused by B. Pertussis

CDC tracks Pertussis # cases and can be accessed here

Remember: to generate the direct link in the word "here", you can use text

Now, how do we scrape the data, if there's not downloadable version of it? One way to do this is by using "datapasta" package.

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.

```
##install.packages("datapasta")
```

Datapasta will add some Addins that serve to specify which type of data we are scrapping.

```
## copy the table data from the website. Go to Addins and hit Paste as data.frame
cdc <- data.frame(
    year = c(
        1922L, 1923L, 1924L, 1925L,
        1926L, 1927L, 1928L, 1929L, 1930L, 1931L,
        1932L, 1933L, 1934L, 1935L, 1936L,
        1937L, 1938L, 1939L, 1940L, 1941L, 1942L,
        1943L, 1944L, 1945L, 1946L, 1947L,
        1948L, 1949L, 1950L, 1951L, 1952L,
        1953L, 1954L, 1955L, 1956L, 1957L, 1958L,
        1959L, 1960L, 1961L, 1962L, 1963L,
        1964L, 1965L, 1966L, 1967L, 1968L, 1969L,
        1970L, 1971L, 1972L, 1973L, 1974L,
        1975L, 1976L, 1977L, 1978L, 1979L, 1980L,</pre>
```

```
1981L, 1982L, 1983L, 1984L, 1985L,
    1986L, 1987L, 1988L, 1989L, 1990L,
    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, 2020L, 2021L, 2022L, 2024L
  ),
  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,
    6124, 2116, 3044, 23544
  )
)
##install.packages("styler"), then apply "style selection"
```

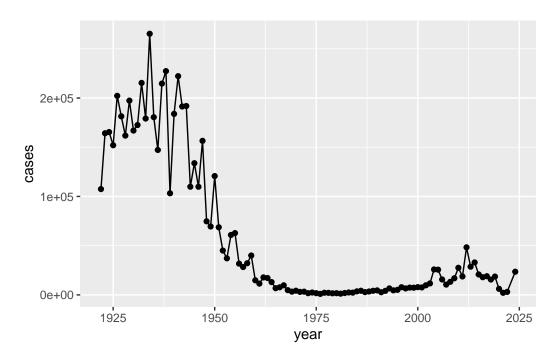
Let's plot year vs. cases to see the trend over time in the US

```
library(ggplot2)
```

```
baseplot <-
    ggplot(cdc) +
    aes(year, cases) +
    geom_point() +</pre>
```

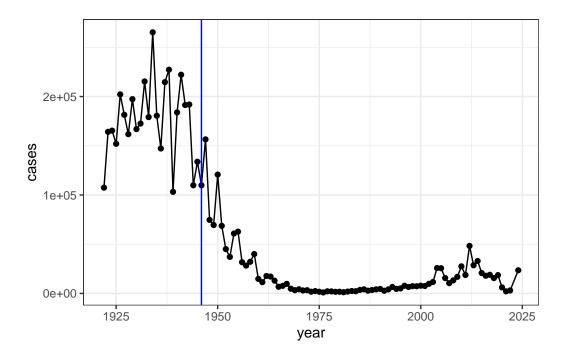
```
geom_line()
```

baseplot



Let's add the date

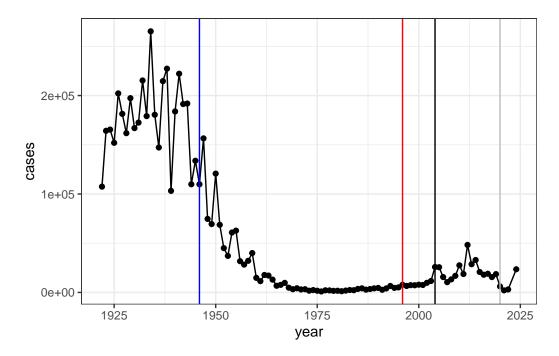
```
baseplot +
  theme_bw() +
  geom_vline(xintercept = 1946, col = "blue")
```



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?

Let's add the date wP vaccine

```
baseplot +
  theme_bw() +
  geom_vline(xintercept = 1946, col = "blue") +
  geom_vline(xintercept = 1996, col = "red") +
  geom_vline(xintercept = 2020, col = "gray") +
  geom_vline(xintercept = 2004)
```



After discussing, we're seeing that the 2nd vax (AP) had a pattern of outbreaks after ~4yrs. So CDC realized this and now the recommendation is every 10yrs for the DTAP vax.

Exploring CMI-PB data. > Q. Why is this vaccine-preventable disease on the upswing?

To answer this question we need to investigate the mechanisms underlying waning protection against pertussis. This requires evaluation of pertussis-specific immune responses over time in wP and aP vaccinated individuals.

##CMI-PB (Computational Models of Immunity - Pertussis Boost) cmi website Very valuable to build predictable models. Data is available to the public. Helps to be a live predictive system.

We can monitor subjects given the 1st or 2nd vax, and their immune system response/gene expression after a period of time (i.e. pre vs post vaccination data)

You can find this really cool project here

Go to Access the data. We're going to pull out this API data, which looks overwhelming - take a look here. But it's simple a format that has an "key item", followed by a "value".

Now to make sense of this API data, we'll use Json to return the values in a analyzable way and make sense of it. (key:value pairs)

We can use **jsonlite** package and it's read_jason() function

```
##install.packages("jsonlite")
library(jsonlite)
subject <- read_json("https://www.cmi-pb.org/api/v5/subject",</pre>
               simplifyVector = TRUE)
Let's have a weep peak
##head(subject)
     Q. How many subjects do we have?
nrow(subject)
[1] 172
     Q. How many male/female?
table(subject$biological_sex)
Female
         Male
           60
   112
     Q. How many wP and aP?
table(subject$infancy_vac)
aP wP
87 85
     Q. break down of biological sex and race
table(subject$race, subject$biological_sex)
```

	${\tt Female}$	Male
American Indian/Alaska Native	0	1
Asian	32	12
Black or African American	2	3
More Than One Race	15	4
Native Hawaiian or Other Pacific Islander	1	1
Unknown or Not Reported	14	7
White	48	32

Q. Does this break down reflect the US population?

```
table(subject$dataset)
```

```
2020_dataset 2021_dataset 2022_dataset 2023_dataset 60 36 22 54
```

```
##head(specimen)
```

```
##head(ab_titer)
```

Now, how do we joint different data points from different tables, i.e., subject, specimen, ab_titer tables?

We merge or "join" them to have all the info we need about a given antibody measurement. We are creating a Meta data to compile based on subject id.

```
library (dplyr)
meta <- inner_join(subject, specimen)</pre>
```

```
subject_id infancy_vac biological_sex
                                                       ethnicity race
1
                       wΡ
                                  Female Not Hispanic or Latino White
2
           1
                       wP
                                  Female Not Hispanic or Latino White
3
           1
                       wP
                                  Female Not Hispanic or Latino White
4
           1
                       wP
                                  Female Not Hispanic or Latino White
5
           1
                       wP
                                  Female Not Hispanic or Latino White
                                  Female Not Hispanic or Latino White
           1
                       wP
 year_of_birth date_of_boost
                                    dataset specimen_id
     1986-01-01
                   2016-09-12 2020_dataset
1
                                                        1
2
     1986-01-01
                    2016-09-12 2020_dataset
                                                       2
3
     1986-01-01
                   2016-09-12 2020_dataset
                                                       3
                                                       4
4
                   2016-09-12 2020_dataset
     1986-01-01
5
     1986-01-01
                   2016-09-12 2020_dataset
                                                       5
                   2016-09-12 2020_dataset
                                                       6
6
     1986-01-01
 actual_day_relative_to_boost planned_day_relative_to_boost specimen_type
                             -3
                                                                        Blood
1
                                                              0
2
                                                                        Blood
                              1
                                                              1
3
                              3
                                                              3
                                                                        Blood
4
                              7
                                                              7
                                                                        Blood
5
                             11
                                                             14
                                                                        Blood
6
                             32
                                                             30
                                                                        Blood
 visit
1
      1
2
      2
3
      3
4
      4
5
      5
      6
6
```

and one last join of ab_titer and meta

```
abdata <- inner_join(meta, ab_titer)</pre>
```

head(abdata)

```
3
                                   Female Not Hispanic or Latino White
           1
                       wP
4
           1
                       wP
                                   Female Not Hispanic or Latino White
5
           1
                       wP
                                   Female Not Hispanic or Latino White
6
           1
                       wP
                                   Female Not Hispanic or Latino White
                                     dataset specimen id
  year_of_birth date_of_boost
                    2016-09-12 2020_dataset
1
     1986-01-01
2
     1986-01-01
                    2016-09-12 2020 dataset
                                                        1
3
     1986-01-01
                    2016-09-12 2020_dataset
                                                        1
                    2016-09-12 2020_dataset
                                                        1
4
     1986-01-01
5
     1986-01-01
                    2016-09-12 2020_dataset
                                                        1
6
                    2016-09-12 2020_dataset
     1986-01-01
                                                        1
  actual_day_relative_to_boost planned_day_relative_to_boost specimen_type
1
                             -3
                                                                         Blood
                             -3
2
                                                              0
                                                                         Blood
3
                             -3
                                                              0
                                                                         Blood
4
                             -3
                                                              0
                                                                         Blood
5
                             -3
                                                              0
                                                                         Blood
                             -3
6
                                                              0
                                                                         Blood
  visit isotype is_antigen_specific antigen
                                                      MFI MFI_normalised unit
1
      1
                               FALSE
                                        Total 1110.21154
                                                                2.493425 UG/ML
            IgE
2
      1
            IgE
                               FALSE
                                        Total 2708.91616
                                                                2.493425 IU/ML
3
      1
            IgG
                                TRUE
                                           PΤ
                                                68.56614
                                                                3.736992 IU/ML
4
      1
            IgG
                                TRUE
                                          PRN 332.12718
                                                                2.602350 IU/ML
5
                                TRUE
                                          FHA 1887.12263
                                                               34.050956 IU/ML
      1
            IgG
6
      1
            IgE
                                TRUE
                                          ACT
                                                 0.10000
                                                                1.000000 IU/ML
  lower_limit_of_detection
1
                   2.096133
2
                  29.170000
3
                   0.530000
4
                   6.205949
5
                   4.679535
6
                   2.816431
```

nrow(abdata)

[1] 52576

table(abdata\$isotype)

```
IgE IgG IgG1 IgG2 IgG3 IgG4
6698 5389 10117 10124 10124 10124
```

table(abdata\$antigen)

ACT	BETV1	DT	FELD1	FHA	FIM2/3	LOLP1	LOS	Measles	OVA
1970	1970	4978	1970	5372	4978	1970	1970	1970	4978
PD1	PRN	PT	PTM	Total	TT				
1970	5372	5372	1970	788	4978				

Let's begin w IgG Filter with dplyr

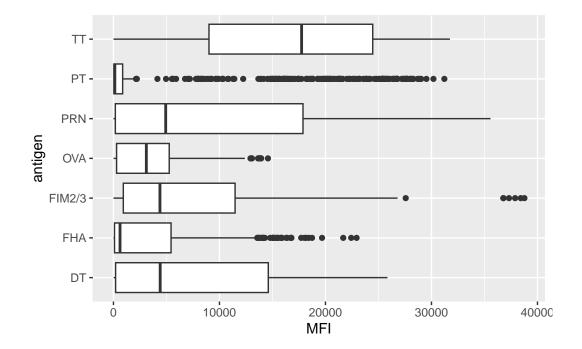
```
igg <- filter(abdata, isotype == "IgG")
head(igg)</pre>
```

```
subject_id infancy_vac biological_sex
                                                       ethnicity race
           1
                      wP
                                  Female Not Hispanic or Latino White
1
2
           1
                       wP
                                  Female Not Hispanic or Latino White
3
           1
                       wP
                                  Female Not Hispanic or Latino White
4
           1
                      wP
                                  Female Not Hispanic or Latino White
5
           1
                                  Female Not Hispanic or Latino White
                       wP
           1
                      wP
                                  Female Not Hispanic or Latino White
  year_of_birth date_of_boost
                                    dataset specimen_id
1
     1986-01-01
                   2016-09-12 2020_dataset
2
                    2016-09-12 2020_dataset
     1986-01-01
                                                       1
3
                    2016-09-12 2020_dataset
                                                       1
     1986-01-01
                                                       2
4
                   2016-09-12 2020_dataset
     1986-01-01
                                                       2
5
     1986-01-01
                    2016-09-12 2020_dataset
     1986-01-01
                   2016-09-12 2020_dataset
                                                       2
  actual_day_relative_to_boost planned_day_relative_to_boost specimen_type
1
                             -3
                                                                        Blood
2
                             -3
                                                             0
                                                                        Blood
3
                             -3
                                                             0
                                                                        Blood
4
                              1
                                                              1
                                                                        Blood
5
                              1
                                                              1
                                                                        Blood
6
                              1
                                                              1
                                                                        Blood
  visit isotype is_antigen_specific antigen
                                                     MFI MFI_normalised unit
      1
                                                68.56614
                                                                3.736992 IU/ML
1
            IgG
                                TRUE
                                          PΤ
2
      1
            IgG
                                TRUE
                                         PRN 332.12718
                                                                2.602350 IU/ML
3
      1
            IgG
                                TRUE
                                         FHA 1887.12263
                                                               34.050956 IU/ML
      2
                                           PΤ
4
                                TRUE
                                                41.38442
                                                                2.255534 IU/ML
            IgG
5
      2
            IgG
                                TRUE
                                         PRN 174.89761
                                                                1.370393 IU/ML
```

```
6
      2
            IgG
                                TRUE
                                          FHA 246.00957
                                                                4.438960 IU/ML
 lower_limit_of_detection
1
                  0.530000
2
                  6.205949
3
                  4.679535
4
                  0.530000
5
                  6.205949
                  4.679535
6
```

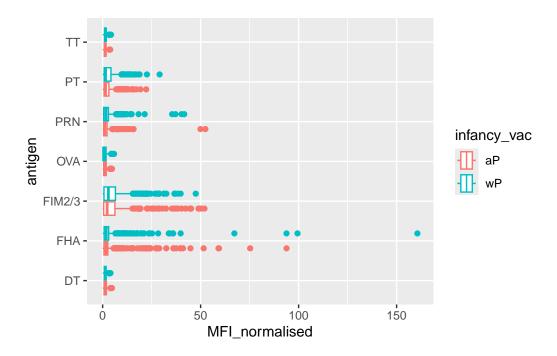
Let's make a box plot of Ig levels – A plot of MFI vs antigen

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



To make a plot by infancy vac differentiating the two vaxs by color

```
ggplot(igg) +
  aes(MFI_normalised, antigen, col=infancy_vac) + ## remember "col" is for coloring
  geom_boxplot()
```



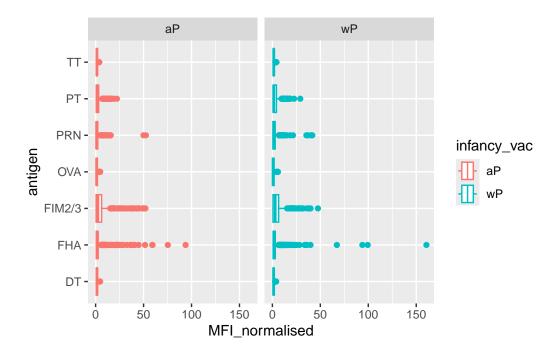
Ideally, I would like to see how these Ab levels change over time relative to the booster shot.

```
table(abdata$visit) ## How many visits?
```

```
1 2 3 4 5 6 7 8 9 10 11 12
8280 8280 8420 6565 6565 6210 5810 815 735 686 105 105
```

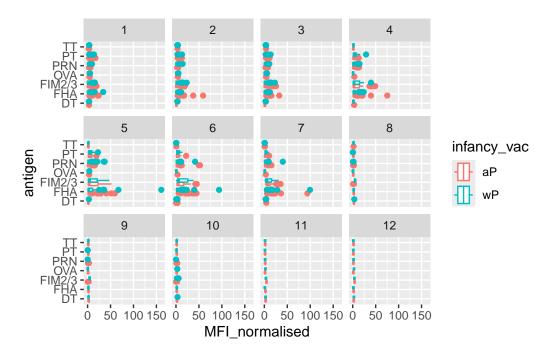
Faceting the data by infancy vac

```
ggplot(igg) +
  aes(MFI_normalised, antigen, col=infancy_vac) +
  geom_boxplot() +
  facet_wrap(~infancy_vac)
```



Faceting the data by visit and differentiated by infancy vax type

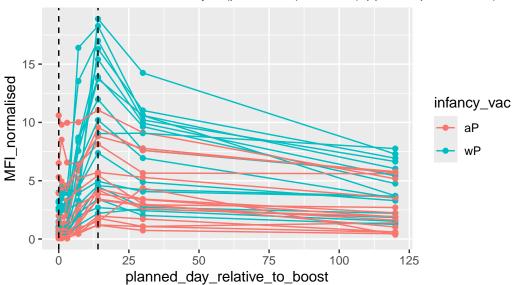
```
ggplot(igg) +
  aes(MFI_normalised, antigen, col=infancy_vac) +
  geom_boxplot() +
  facet_wrap(~visit)
```



Make a line plot of 2021 data using dataset IgG PT. Where each line of teh plot is a subject

2021 dataset IgG PT

Dashed lines indicate day 0 (pre-boost) and 14 (apparent peak levels)

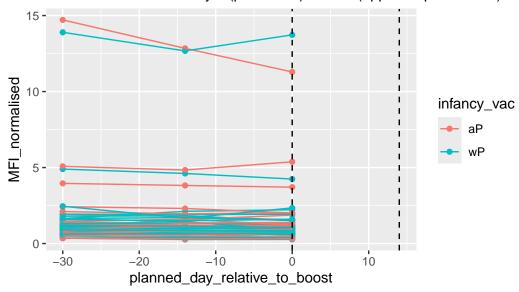


```
abdata.23 <- abdata %>% filter(dataset == "2023_dataset")

abdata.23 %>%
filter(isotype == "IgG", antigen == "PT") %>%
ggplot() +
    aes(x=planned_day_relative_to_boost,
        y=MFI_normalised,
        col=infancy_vac,
        group=subject_id) +
    geom_point() +
    geom_point() +
    geom_vline(xintercept=0, linetype="dashed") +
    geom_vline(xintercept=14, linetype="dashed") +
    labs(title="2023 dataset IgG PT",
        subtitle = "Dashed lines indicate day 0 (pre-boost) and 14 (apparent peak levels)")
```

2023 dataset IgG PT

Dashed lines indicate day 0 (pre-boost) and 14 (apparent peak levels)



Obtaining CMI-PB RNASeq data, see here