lab19

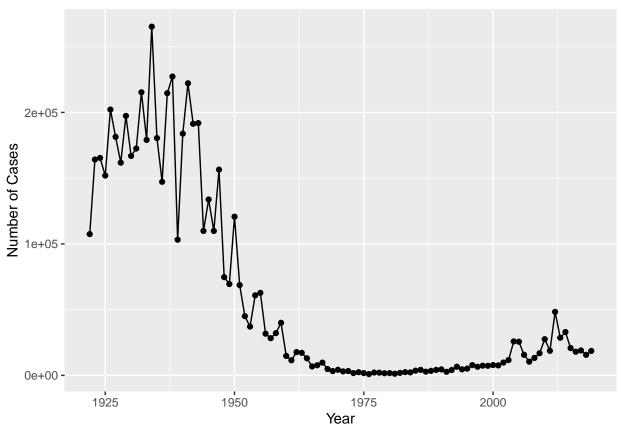
Zijing

2022-12-02

Q1

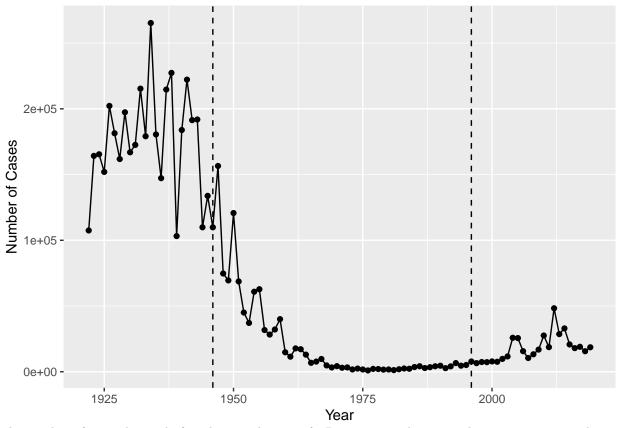
```
cdc <- data.frame(</pre>
                           Year = c(1922L,
                                     1923L,1924L,1925L,1926L,1927L,1928L,
                                     1929L,1930L,1931L,1932L,1933L,1934L,1935L,
                                     1936L,1937L,1938L,1939L,1940L,1941L,
                                     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,
                                     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),
  No..Reported.Pertussis.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)
library(ggplot2)
ggplot(cdc) +
 aes(Year, No..Reported.Pertussis.Cases) +
```

```
geom_point() +
geom_line() +
labs(x="Year",y="Number of Cases")
```



```
# Q2
```

```
ggplot(cdc) +
  aes(Year, No..Reported.Pertussis.Cases) +
  geom_point() +
  geom_line() +
  geom_vline(xintercept=c(1946,1996),linetype="dashed")+
  labs(x="Year",y="Number of Cases")
```



The number of cases dropped after the introduction of wP vaccination but started to rise again several years after the introduction of aP vaccination.

$\mathbf{Q3}$

The number of cases started to rise again several years after the introduction of aP vaccination. Maybe this is caused by the infection happening in grown-up infants that got aP vaccination?

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

```
subject_id infancy_vac biological_sex
##
                                                          ethnicity race
## 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
## 1
        1986-01-01
                       2016-09-12 2020_dataset
## 2
        1968-01-01
                       2019-01-28 2020_dataset
## 3
        1983-01-01
                       2016-10-10 2020_dataset
```

$\mathbf{Q4}$

```
table(subject$infancy_vac)
##
## aP wP
## 47 49
47~\mathrm{aP} and 49~\mathrm{wP}.
Q_5
table(subject$biological_sex)
## Female
            Male
       66
66 Female and 30 Male.
table(subject$race)
##
##
                American Indian/Alaska Native
##
##
                                         Asian
##
                                             27
                    Black or African American
##
##
##
                           More Than One Race
##
## Native Hawaiian or Other Pacific Islander
##
                      Unknown or Not Reported
##
##
                                         White
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
```

```
male <- subject%>%filter(biological_sex == "Male")
female <- subject%>%filter(biological_sex == "Female")
Male race breakdown:
table(male$race)
##
                American Indian/Alaska Native
##
##
##
                                         Asian
##
##
                           More Than One Race
##
## Native Hawaiian or Other Pacific Islander
##
                      Unknown or Not Reported
##
##
                                             4
##
                                         White
##
                                            13
Female race breakdown:
table(female$race)
##
##
                                         Asian
##
##
                    Black or African American
##
##
                           More Than One Race
##
## Native Hawaiian or Other Pacific Islander
##
                      Unknown or Not Reported
##
                                            10
##
                                         White
                                            27
library(lubridate)
## Loading required package: timechange
## Attaching package: 'lubridate'
## The following objects are masked from 'package:base':
##
##
       date, intersect, setdiff, union
today()
## [1] "2022-12-03"
today() - ymd("2000-01-01")
```

Time difference of 8372 days

```
time_length( today() - ymd("2000-01-01"), "years")
## [1] 22.92129
```

```
subject$age <- today() - ymd(subject$year_of_birth)
ap <- subject%>%filter(infancy_vac == "aP")
wp <- subject%>%filter(infancy_vac == "wP")
time_length(mean(ap$age), "years")

## [1] 25.23908
time_length(mean(wp$age), "years")
```

[1] 36.08353

wP average: 36 years, aP average: 25 years. The average age of wP receivers is much higher than that of aP receivers

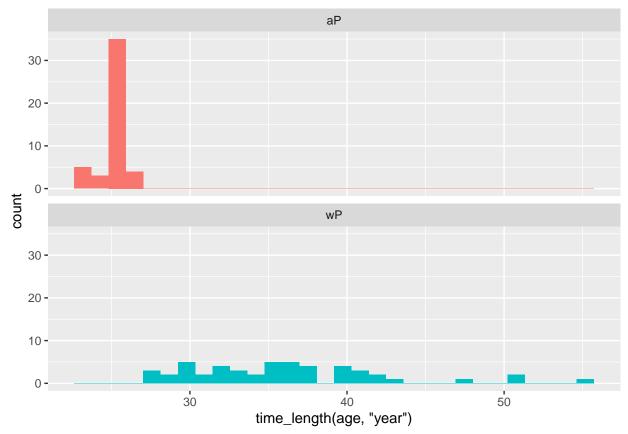
Q8

```
boost_age <- time_length(ymd(subject$date_of_boost) - ymd(subject$year_of_birth), "year")
head(boost_age)
## [1] 30.69678 51.07461 33.77413 28.65982 25.65914 28.77481</pre>
```

$\mathbf{Q}9$

```
ggplot(subject) +
  aes(time_length(age, "year"),
      fill=as.factor(infancy_vac)) +
  geom_histogram(show.legend=FALSE) +
  facet_wrap(vars(infancy_vac), nrow=2)
```

`stat_bin()` using `bins = 30`. Pick better value with `binwidth`.



Yes, these two groups are different.

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

$\mathbf{Q}9$

```
meta <- full_join(specimen, subject)</pre>
## Joining, by = "subject_id"
dim(meta)
## [1] 729
head(meta)
     specimen_id subject_id actual_day_relative_to_boost
##
## 1
                                                          -3
                1
## 2
                2
                                                         736
                            1
                3
## 3
                            1
                                                           1
## 4
                4
                            1
                                                           3
                5
                                                           7
## 5
                            1
## 6
                6
                            1
                                                          11
##
     planned_day_relative_to_boost specimen_type visit infancy_vac biological_sex
## 1
                                   0
                                                                                Female
                                              Blood
                                                         1
                                                                     wΡ
## 2
                                 736
                                              Blood
                                                        10
                                                                     wP
                                                                                Female
## 3
                                              Blood
                                                         2
                                   1
                                                                     wP
                                                                                Female
```

```
## 4
                                            Blood
                                                                  wΡ
                                                                             Female
## 5
                                  7
                                            Blood
                                                      4
                                                                             Female
                                                                  wΡ
## 6
                                 14
                                            Blood
                                                      5
                                                                  wΡ
                                                                             Female
##
                  ethnicity race year_of_birth date_of_boost
                                                                     dataset
## 1 Not Hispanic or Latino White
                                      1986-01-01
                                                    2016-09-12 2020_dataset
                                                    2016-09-12 2020 dataset
## 2 Not Hispanic or Latino White
                                      1986-01-01
## 3 Not Hispanic or Latino White
                                      1986-01-01
                                                    2016-09-12 2020 dataset
## 4 Not Hispanic or Latino White
                                      1986-01-01
                                                    2016-09-12 2020_dataset
## 5 Not Hispanic or Latino White
                                      1986-01-01
                                                    2016-09-12 2020 dataset
## 6 Not Hispanic or Latino White
                                      1986-01-01
                                                    2016-09-12 2020_dataset
            age
## 1 13485 days
## 2 13485 days
## 3 13485 days
## 4 13485 days
## 5 13485 days
## 6 13485 days
```

```
abdata <- inner_join(titer, meta)</pre>
## Joining, by = "specimen_id"
dim(abdata)
## [1] 32675
                 21
head(abdata)
     specimen_id isotype is_antigen_specific antigen
##
                                                                MFI MFI_normalised
## 1
               1
                                         FALSE
                                                  Total 1110.21154
                      IgE
## 2
                1
                                         FALSE
                                                  Total 2708.91616
                                                                          2.493425
                      IgE
## 3
                1
                      IgG
                                          TRUE
                                                     PT
                                                          68.56614
                                                                          3.736992
## 4
                                          TRUE
                                                    PRN
                1
                      IgG
                                                         332.12718
                                                                          2.602350
               1
                      IgG
                                          TRUE
                                                    FHA 1887.12263
                                                                         34.050956
                                                    ACT
## 6
                                          TRUE
                                                            0.10000
                1
                      IgE
                                                                           1.000000
      unit lower_limit_of_detection subject_id actual_day_relative_to_boost
##
## 1 UG/ML
                            2.096133
                                                                              -3
## 2 IU/ML
                           29.170000
                                                                              -3
                                                                              -3
## 3 IU/ML
                            0.530000
                                                1
## 4 IU/ML
                            6.205949
                                                                              -3
## 5 IU/ML
                                                1
                                                                              -3
                            4.679535
## 6 IU/ML
                            2.816431
                                                1
##
     planned_day_relative_to_boost specimen_type visit infancy_vac biological_sex
## 1
                                   0
                                             Blood
                                                        1
                                                                    wΡ
                                                                                Female
## 2
                                   0
                                             Blood
                                                        1
                                                                    wΡ
                                                                                Female
## 3
                                   0
                                                                    wP
                                                                                Female
                                             Blood
                                                        1
                                   0
## 4
                                             Blood
                                                                    wP
                                                                                Female
## 5
                                   0
                                             Blood
                                                        1
                                                                    wP
                                                                                Female
## 6
                                   0
                                                                    wP
                                                                                Female
                                             Blood
##
                   ethnicity race year_of_birth date_of_boost
                                                                       dataset
## 1 Not Hispanic or Latino White
                                       1986-01-01
                                                      2016-09-12 2020_dataset
## 2 Not Hispanic or Latino White
                                       1986-01-01
                                                      2016-09-12 2020_dataset
```

```
## 3 Not Hispanic or Latino White
                                     1986-01-01
                                                    2016-09-12 2020_dataset
                                                    2016-09-12 2020_dataset
## 4 Not Hispanic or Latino White
                                     1986-01-01
## 5 Not Hispanic or Latino White
                                     1986-01-01
                                                    2016-09-12 2020 dataset
                                                    2016-09-12 2020_dataset
## 6 Not Hispanic or Latino White
                                     1986-01-01
            age
## 1 13485 days
## 2 13485 days
## 3 13485 days
## 4 13485 days
## 5 13485 days
## 6 13485 days
```

```
table(abdata$isotype)
```

Q12

table(abdata\$visit)

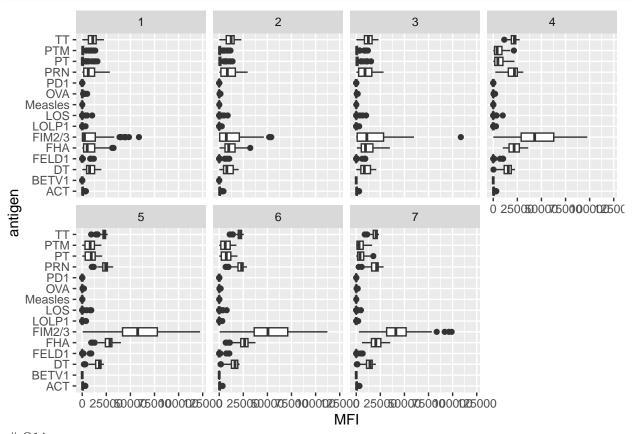
Much less specimens in visit 8 compared to other visits.

```
ig1 <- abdata %>% filter(isotype == "IgG1", visit!=8)
head(ig1)
```

```
specimen_id isotype is_antigen_specific antigen
                                                               MFI MFI_normalised
## 1
                                          TRUE
                                                   ACT 274.355068
                                                                         0.6928058
               1
                     IgG1
                                          TRUE
                                                   LOS 10.974026
                                                                         2.1645083
               1
                     IgG1
## 3
                     IgG1
                                          TRUE
                                                 FELD1
                                                          1.448796
                                                                         0.8080941
               1
## 4
                                          TRUE
                                                 BETV1
                                                          0.100000
                                                                         1.0000000
               1
                     IgG1
## 5
               1
                     IgG1
                                          TRUE
                                                 LOLP1
                                                          0.100000
                                                                         1.0000000
               1
                     IgG1
                                          TRUE Measles 36.277417
                                                                         1.6638332
      unit lower_limit_of_detection subject_id actual_day_relative_to_boost
##
## 1 IU/ML
                            3.848750
                                               1
                                                                             -3
## 2 IU/ML
                                               1
                                                                             -3
                            4.357917
                                                                             -3
## 3 IU/ML
                            2.699944
                                               1
## 4 IU/ML
                            1.734784
                                               1
                                                                             -3
## 5 IU/ML
                            2.550606
                                               1
                                                                             -3
## 6 IU/ML
                            4.438966
                                               1
     planned_day_relative_to_boost specimen_type visit infancy_vac biological_sex
## 1
                                  0
                                             Blood
                                                        1
                                                                   wP
                                                                               Female
                                                                   wP
## 2
                                  0
                                                                               Female
                                             Blood
                                                        1
## 3
                                  0
                                             Blood
                                                        1
                                                                   wP
                                                                               Female
## 4
                                  0
                                             Blood
                                                                   wP
                                                                               Female
                                                        1
```

```
## 5
                                            Blood
                                                                  wP
                                                                             Female
## 6
                                                      1
                                  0
                                            Blood
                                                                  wP
                                                                             Female
##
                                                                     dataset
                  ethnicity race year_of_birth date_of_boost
## 1 Not Hispanic or Latino White
                                      1986-01-01
                                                     2016-09-12 2020_dataset
## 2 Not Hispanic or Latino White
                                      1986-01-01
                                                     2016-09-12 2020_dataset
## 3 Not Hispanic or Latino White
                                      1986-01-01
                                                     2016-09-12 2020 dataset
## 4 Not Hispanic or Latino White
                                      1986-01-01
                                                     2016-09-12 2020 dataset
                                                     2016-09-12 2020_dataset
## 5 Not Hispanic or Latino White
                                      1986-01-01
## 6 Not Hispanic or Latino White
                                      1986-01-01
                                                     2016-09-12 2020_dataset
##
            age
## 1 13485 days
## 2 13485 days
## 3 13485 days
## 4 13485 days
## 5 13485 days
## 6 13485 days
```

```
ggplot(ig1) +
aes(MFI, antigen) +
geom_boxplot() +
facet_wrap(vars(visit), nrow=2)
```

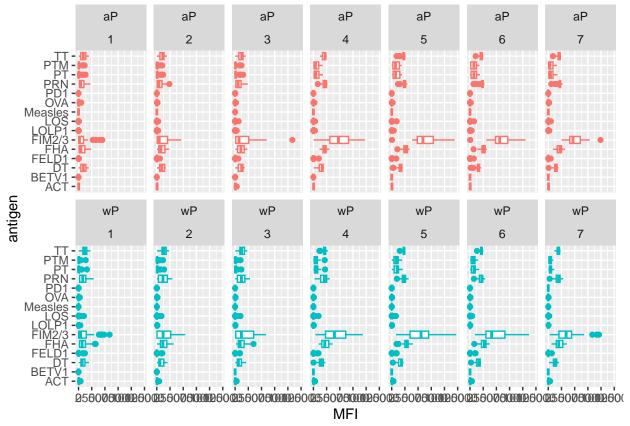


Q14

FIM2/3 is the most different one antigen. PRN and FHA are quite different as well.

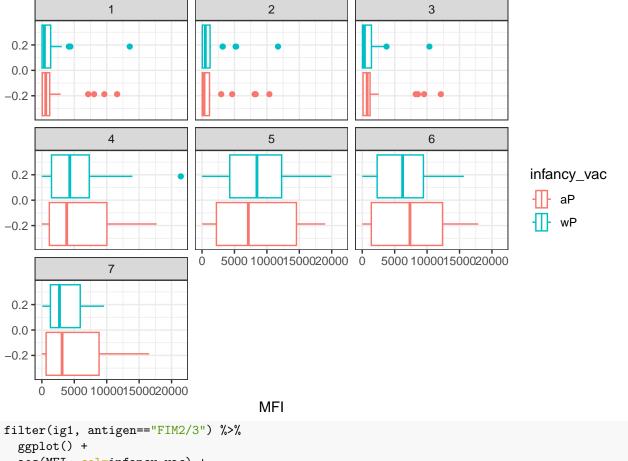
```
ggplot(ig1) +
 aes(MFI, antigen, col=infancy_vac ) +
 geom_boxplot(show.legend = FALSE) +
 facet_wrap(vars(visit), nrow=2) +
 theme_bw()
                               2
                                               3
                                                               4
               1
    PŔŃ
    PD1
OVA
  Measles
   LOS
LOLP1
FIM2/3
antigen
                                                         5
                               6
                                               7
    OVA
  Measles
   FHA
FELD1
     DT
        MFI
```

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

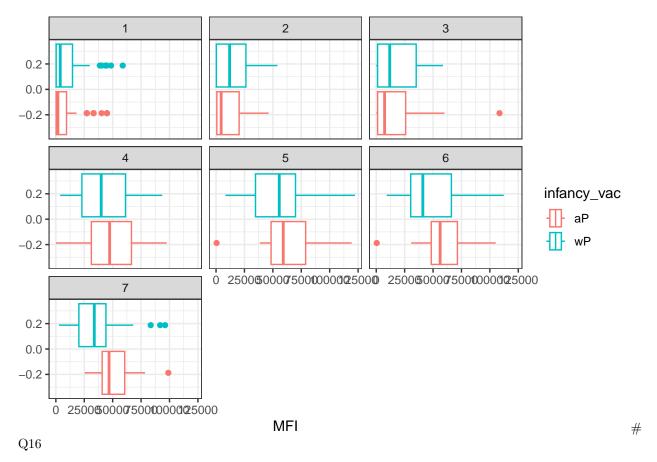


```
# Q15
```

```
filter(ig1, antigen=="PTM") %>%
    ggplot() +
    aes(MFI, col=infancy_vac) +
    geom_boxplot(show.legend = TRUE) +
    facet_wrap(vars(visit)) +
    theme_bw()
```



```
filter(ig1, antigen=="FIM2/3") %>%
    ggplot() +
    aes(MFI, col=infancy_vac) +
    geom_boxplot(show.legend = TRUE) +
    facet_wrap(vars(visit)) +
    theme_bw()
```



Both rise over time, but FIM2/3 more significantly. Visit 5 seems to be the peak for both.

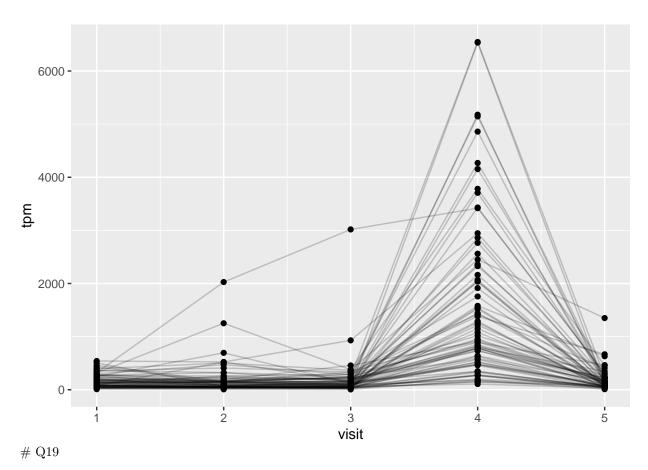
Q17

aP response starts being lower but ends up being higher than wP response. However, they follow a similar trend of rising and then declining over time.

```
url <- "https://www.cmi-pb.org/api/v2/rnaseq?versioned_ensembl_gene_id=eq.ENSG00000211896.7"
rna <- read_json(url, simplifyVector = TRUE)
ssrna <- inner_join(rna, meta)
## Joining, by = "specimen_id"</pre>
```

Q18

```
ggplot(ssrna) +
  aes(visit, tpm, group=subject_id) +
  geom_point() +
  geom_line(alpha=0.2)
```



The gene expression is at its maximum level at visit 4.

$\mathbf{Q20}$

This does not match the antibody pattern as the antibody peaks at visit 5. This is likely because antibodies are made after the genes are expressed and would live for a long time. Thus, the antibody expression continues to accumulate until gene expression has dropped to zero, which is until sometime between visit 4 and 5, leading to the peak in antibody detection on visit 5.

```
ggplot(ssrna) +
aes(tpm, col=infancy_vac) +
geom_boxplot() +
facet_wrap(vars(visit))
```

