

Race

```
library(data.table)
library(dplyr)

##
## Attaching package: 'dplyr'

## The following objects are masked from 'package:data.table':
##
##   between, first, last

## The following objects are masked from 'package:stats':
##
##   filter, lag

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

library(ggplot2)
library(reshape2)

##
## Attaching package: 'reshape2'

## The following objects are masked from 'package:data.table':
##
##   dcast, melt

library(scales)
library(parallel)
library(stringr)
library(knitr)

Load metadata

dt.hip.stats = fread("annotations/hip_stats.txt") %>%
  mutate(race = ifelse(is.na(race), "Unknown", race)) %>%
  mutate(count_total = count, occurrences_total = diversity) %>%
  select(sample_id, race)

# split race and origin

tmp = str_split_fixed(dt.hip.stats$race, ",", n = 2)
tmp[,2] = ifelse(tmp[,2] == "", tmp[,1], tmp[,2])
tmp[,1] = ifelse(tmp[,2] == tmp[,1], "Unknown", tmp[,1])
dt.hip.stats$race = tmp[,1]

dt.hip.stats = dt.hip.stats %>% filter(race != "Unknown")

summary(as.factor(dt.hip.stats$race))

## asian or pacific islander black or african american
##               47               12
##      caucasian
##             465
```

Load VDJdb annotations with 1 mismatch for HIP data (time consuming, ~ 2mln clonotypes)

```
dt.hip = rbindlist(mclapply(as.list(dt.hip.stats$sample_id),
                           function(x) fread(paste0("annotations/split_1mm/", x, ".annot.txt")) %>%
                             mutate(sample_id = x), mc.cores = 40)) %>%
  group_by(sample_id, cdr3) %>%
  summarise(count = sum(count), occurrences = n())
```

VDJdb data

```
dt.vdjdbc = fread("rearr_model/VDJDB_fullP_rob_ageing.txt") %>%
  filter(gene == "TRB", mhc.class == "MHCI") %>%
  mutate(hla_spec = str_split_fixed(mhc.a, pattern = "[:,]", 2)[,1]) %>%
  select(cdr3, hla_spec, antigen.epitope, antigen.species) %>%
  group_by(antigen.epitope) %>%
  mutate(unique_cdrs = n()) %>%
  filter(unique_cdrs > 30) %>%
  select(cdr3, antigen.epitope, antigen.species, unique_cdrs)
```

Merge

```
dt.hip.m = dt.hip %>%
  merge(dt.hip.stats) %>%
  merge(dt.vdjdbc)
```

Summarise by race

```
dt.hip.s = as.data.table(dt.hip.m) %>%
  group_by(sample_id, race, antigen.epitope, antigen.species, unique_cdrs) %>%
  summarise(occurrences = sum(occurrences)) %>%
  group_by(sample_id) %>%
  mutate(occurrences_share = occurrences / sum(occurrences) / unique_cdrs)
```

```
dt.p = data.table(antigen.epitope = unique(dt.hip.s$antigen.epitope), p = 1, freq.ratio_c = 1, freq.ratio_b = 1, freq.ratio_a = 1)
merge(dt.hip.s %>% ungroup %>% select(antigen.species, antigen.epitope) %>% unique)
```

```
for (i in 1:nrow(dt.p)) {
  tmp = dt.hip.s %>% filter(antigen.epitope == dt.p$antigen.epitope[i])
  tmp$race = as.factor(tmp$race)
  dt.p$freq.ratio_c[i] = with(tmp, mean(occurrences_share[which(race=="caucasian")])) / mean(occurrences_share)
  dt.p$freq.ratio_b[i] = with(tmp, mean(occurrences_share[which(race=="black or african american")])) / mean(occurrences_share)
  dt.p$freq.ratio_a[i] = with(tmp, mean(occurrences_share[which(race=="asian or pacific islander")])) / mean(occurrences_share)
  a = aov(occurrences_share ~ race, tmp)
  dt.p$p[i] = summary(a)[[1]][["Pr(>F)"]][1]
  #dt.p$p[i] = kruskal.test(occurrences_share ~ race, tmp)$p.value
}
```

```
dt.p$p = p.adjust(dt.p$p, method = "BH")
dt.p$len = nchar(as.character(dt.p$antigen.epitope))
```

```
kable(dt.p %>% arrange(p))
```

| antigen.epitope | p | freq.ratio_c | freq.ratio_b | freq.ratio_a | antigen.species | len |
|-----------------|-----------|--------------|--------------|--------------|-----------------|-----|
| GPGHKARVL | 0.0031257 | 0.9928759 | 0.8924625 | 1.0979395 | HIV-1 | 9 |
| KLVALGINAV | 0.0031257 | 0.9941108 | 1.0160172 | 1.0541764 | HCV | 10 |
| KRWIILGLNK | 0.0031257 | 0.9960243 | 1.0080219 | 1.0372860 | HIV-1 | 10 |

| antigen.epitope | p | freq.ratio_c | freq.ratio_b | freq.ratio_a | antigen.species | len |
|-----------------|-----------|--------------|--------------|--------------|------------------|-----|
| TPRVTGGGAM | 0.0031257 | 1.0086070 | 0.9828783 | 0.9192171 | CMV | 10 |
| KAFSPEVIPMF | 0.0062588 | 1.0065380 | 0.9696819 | 0.9430560 | HIV-1 | 11 |
| EIYKRWH | 0.0091940 | 0.9898537 | 1.1061235 | 1.0732886 | HIV-1 | 8 |
| HSKKKCDEL | 0.0091940 | 1.0181298 | 0.9496281 | 0.8342629 | HCV | 9 |
| VTEHDTLLY | 0.0115844 | 0.9977734 | 0.9667804 | 1.0305104 | CMV | 9 |
| LPPIVAKEI | 0.0157459 | 0.9934615 | 1.1069874 | 1.0373736 | HIV-1 | 9 |
| FLYNLLTRV | 0.0737289 | 1.0047088 | 0.9742584 | 0.9599855 | HomoSapiens | 9 |
| LPRRSGAAGA | 0.1541243 | 0.9983015 | 0.9824625 | 1.0212824 | InfluenzaA | 10 |
| CINGVCWTV | 0.1554756 | 1.0062263 | 0.9397990 | 0.9537703 | HCV | 9 |
| ISPRTLNAW | 0.1554756 | 0.9941388 | 1.0751480 | 1.0388022 | HIV-1 | 9 |
| NLVPMTATV | 0.1727873 | 1.0020015 | 0.9835863 | 0.9843888 | CMV | 9 |
| FPRPWLHGL | 0.2273192 | 1.0035115 | 1.0445965 | 0.9539470 | HIV-1 | 9 |
| KRWIIMGLNK | 0.3251211 | 0.9951243 | 1.0289588 | 1.0408442 | HIV-1 | 10 |
| GILGFVFTL | 0.3965058 | 1.0016895 | 1.0024012 | 0.9826721 | InfluenzaA | 9 |
| LLLIGILV | 0.4854929 | 0.9985625 | 1.0167356 | 1.0099488 | HomoSapiens | 9 |
| AMFWSVPTV | 0.7671592 | 0.9976709 | 1.0112606 | 1.0201682 | HomoSapiens | 9 |
| ATDALMTGY | 0.9484594 | 1.0022560 | 0.9810788 | 0.9825108 | HCV | 9 |
| EPLPQGQLTAY | 0.9557428 | 1.0033693 | 1.0138007 | 0.9632132 | EBV | 11 |
| FLKEKGGL | 0.9557428 | 0.9994419 | 1.0208198 | 1.0002058 | HIV-1 | 8 |
| GLCTLVAML | 0.9557428 | 0.9993633 | 1.0113999 | 1.0033884 | EBV | 9 |
| HPKVSSEVHI | 0.9557428 | 1.0015663 | 1.0152859 | 0.9806006 | HIV-1 | 10 |
| IIKDYGKQM | 0.9557428 | 1.0003727 | 1.0325668 | 0.9879974 | HIV-1 | 9 |
| IPSINVHHY | 0.9557428 | 0.9991849 | 0.9819760 | 1.0126657 | CMV | 9 |
| LLWNGPMAV | 0.9557428 | 0.9992480 | 1.0128237 | 1.0041655 | YellowFeverVirus | 9 |
| RAKFKQLL | 0.9557428 | 1.0009982 | 0.9847979 | 0.9940055 | EBV | 8 |
| RPRGEVRFL | 0.9557428 | 1.0008575 | 1.0490814 | 0.9790218 | HSV-2 | 9 |
| TPGPGVRYPL | 0.9557428 | 1.0016117 | 0.9961808 | 0.9850295 | HIV-1 | 10 |
| YVLDHLIVV | 0.9557428 | 0.9987287 | 1.0175173 | 1.0081052 | EBV | 9 |
| ELAGIGILTV | 0.9870871 | 0.9999765 | 1.0013629 | 0.9998846 | HomoSapiens | 10 |
| SLYNTVATL | 0.9870871 | 1.0000705 | 1.0100585 | 0.9967363 | HIV-1 | 9 |
| TPQDLNTML | 0.9870871 | 1.0002085 | 1.0046113 | 0.9967597 | HIV-1 | 9 |

```

good_epi = (dt.p %>% filter(p < 0.05))$antigen.epitope
dt.hip.s = dt.hip.s %>%
  mutate(antigen.epitope = ifelse(antigen.epitope %in% good_epi, paste(antigen.epitope, "(*)"), antigen.epitope))

dt.hip.s.s = dt.hip.s %>%
  #filter(ucb == T) %>%
  group_by(antigen.epitope) %>%
  summarise(freq = mean(occurrences_share[which(race == "caucasian"])))

dt.hip.s$antigen.epitope = factor(dt.hip.s$antigen.epitope,
  levels = dt.hip.s.s$antigen.epitope[order(dt.hip.s.s$freq)])

dt.hip.s$race = toupper(substr(as.character(dt.hip.s$race), 1, 1))

p19=ggplot(dt.hip.s, aes(x = antigen.epitope, group = paste(antigen.epitope,race),
  fill = race,
  y = occurrences_share)) +
  geom_boxplot(color = "black") +
  coord_flip() +
  scale_fill_brewer(palette = "Set1") +

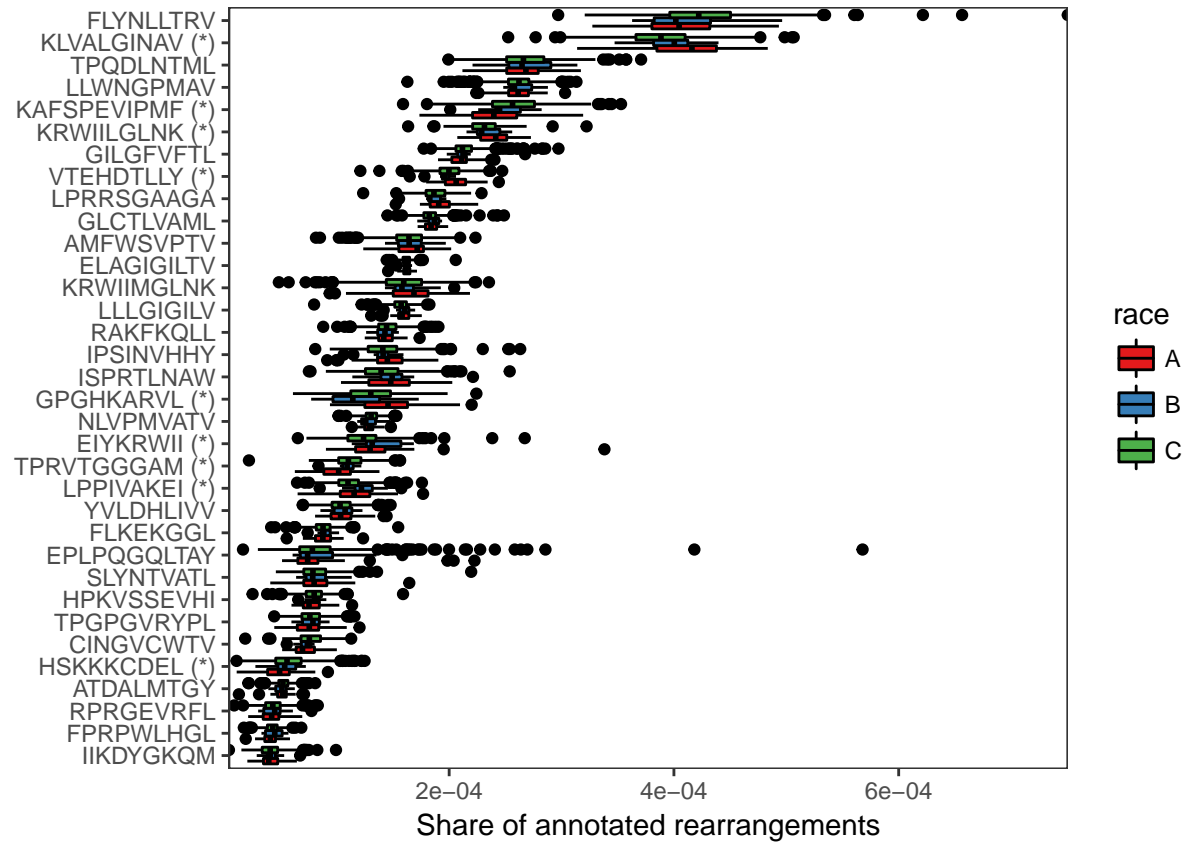
```

```

scale_color_brewer(palette = "Set1") +
xlab("") + scale_y_continuous("Share of annotated rearrangements",
                             expand = c(0,0)) +

theme_bw() +
theme(panel.grid.major = element_blank(), panel.grid.minor = element_blank())
p19

```



```

ggsave("figures/p19.pdf", p19)

```

```
## Saving 6.5 x 4.5 in image
```