

# Results Section: Public Genetic Diversity

```
library(staphopia)
library(ggplot2)
library(reshape2)
library(scales)
USE_DEV = TRUE
```

In this section we will look into genetic diversity that has been sequenced in *Staphylococcus aureus*. In order to do so, we'll use variant counts, cgMLST and MLST as measures of diversity.

## Aggregating Data For Public Samples

First we'll get all publicly available *S. aureus* samples.

```
ps <- get_public_samples()
```

## MLST

Next we will use the MLST information as a measure of genetic diversity. In this case we are interested in the total number of unique sequence types sequenced. We'll use `get_st_by_year()` to get some basic stats about how many STs have been sequenced. We will also use `get_top_sequence_types()` to get each ST represented in the database and the total number of samples with each ST. (*Note: 5000 is just an arbitrarily large number to retrieve all STs*)

```
sequence_types <- get_st_by_year()
top_st <- get_top_sequence_types(5000)
colnames(sequence_types)
```

```
## [1] "year"                "unique"
## [3] "novel"               "assigned"
## [5] "assigned_agree"      "assigned_disagree"
## [7] "unassigned"          "unassigned_agree"
## [9] "unassigned_disagree" "predicted_novel"
## [11] "all"                 "partial"
## [13] "ariba_blast"         "mentalist_blast"
## [15] "mentalist_ariba"     "single"
## [17] "ariba"               "mentalist"
## [19] "blast"               "count"
## [21] "overall_novel"       "overall_assigned"
## [23] "overall_assigned_agree" "overall_assigned_disagree"
## [25] "overall_unassigned"  "overall_unassigned_agree"
## [27] "overall_unassigned_disagree" "overall_predicted_novel"
## [29] "overall_all"         "overall_partial"
## [31] "overall_ariba_blast" "overall_mentalist_blast"
## [33] "overall_mentalist_ariba" "overall_single"
## [35] "overall_ariba"       "overall_mentalist"
## [37] "overall_blast"       "overall"
```

This gives us 38 columns for each year. These columns are:

1. year: The year.
2. unique: The Number of unique STs for a given year.

3. novel: Number of STs not sequenced previously.
4. assigned: Samples which a ST was determined.
5. assigned\_agree: Samples in which each program that called an ST agreed in ST.
6. assigned\_disagree: Samples in which programs did not each call the same ST.
7. unassigned: Samples which a ST was not determined.
8. unassigned\_agree: Each program was unable to assign an ST.
9. unassigned\_disagree: Samples in which no ST was determined, but each program does not agree
10. predicted\_novel: Samples with a match to each Loci, but allele pattern does not exist.
11. all: Samples with an ST determined with agreement between each program.
12. partial: Samples with an ST determined with agreement between two programs. 13: ariba\_blast: Samples with an ST determined with agreement between Ariba and BLAST.
13. mentalist\_blast: Samples with an ST determined with agreement between MentaLiST and BLAST.
14. mentalist\_ariba: Samples with an ST determined with agreement between MentaLiST and Ariba.
15. single: Samples with an ST determined by only a single program.
16. ariba: Samples with an ST determined by only Ariba.
17. mentalist: Samples with an ST determined by only MentaLiST.
18. blast: Samples with an ST determined by only BLAST.
19. count: Total number of samples in a given year. 21-38: overall\_X: The cumulative totals of previous years for column *x*

## Summary of MLST Diversity

### Assignment Breakdown

```
t(sequence_types[sequence_types$year == max(sequence_types$year),21:38])
```

```
##                                8
## overall_novel                  1098
## overall_assigned               42337
## overall_assigned_agree        42243
## overall_assigned_disagree      94
## overall_unassigned            612
## overall_unassigned_agree       612
## overall_unassigned_disagree    0
## overall_predicted_novel        306
## overall_all                   41226
## overall_partial                922
## overall_ariba_blast            81
## overall_mentalist_blast        669
## overall_mentalist_ariba        172
## overall_single                189
## overall_ariba                  29
## overall_mentalist              111
## overall_blast                  49
## overall                       42949
```

### Top STs

```
top_st[1:10,]
```

```
##      st count percent overall
## 1    22  7189   16.74   16.74
## 2     8  6184   14.40   31.14
## 3     5  4664   10.86   42.00
```

```
## 4  239  3123    7.27   49.27
## 5  398  2326    5.42   54.68
## 6   30  1872    4.36   59.04
## 7   45  1663    3.87   62.91
## 8   15  1172    2.73   65.64
## 9   36   857    2.00   67.64
## 10 105   857    2.00   69.63
```

This gives us 4 columns for each ST, in descending order based on the *count* column. In other words the most represented STs are seen first. These columns are:

1. st: The sequence type.
2. count: The number of samples with given ST.
3. percent: The percent of samples represented by given ST.
4. overall: The percent of samples represented by given ST and previous STs.

How many unique STs represented?

```
nrow(top_st[top_st$st > 0,])
```

```
## [1] 1098
```

How many STs represented by a single sample?

```
nrow(top_st[top_st$count == 1, ])
```

```
## [1] 588
```

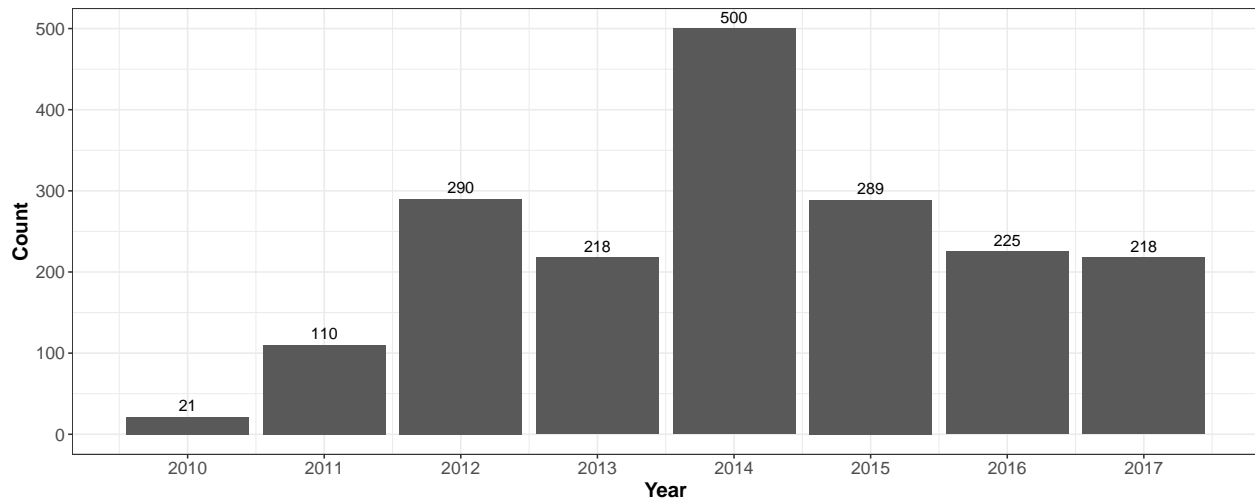
## Visualizing MLST Diversity

The following sections will be plots to visualize relationships in the data.

### Unique Sequence Types By Year

```
p <- ggplot(data=sequence_types, aes(x=year, y=unique)) +
  xlab("Year") +
  ylab("Count") +
  geom_bar(stat='identity') +
  geom_text(aes(label=unique), vjust = -0.5) +
  scale_x_continuous(breaks = round(seq(min(sequence_types$year), max(sequence_types$year), by = 1), 1),
  theme_bw() +
  theme(axis.text=element_text(size=12),
        axis.title=element_text(size=14,face="bold"))
```

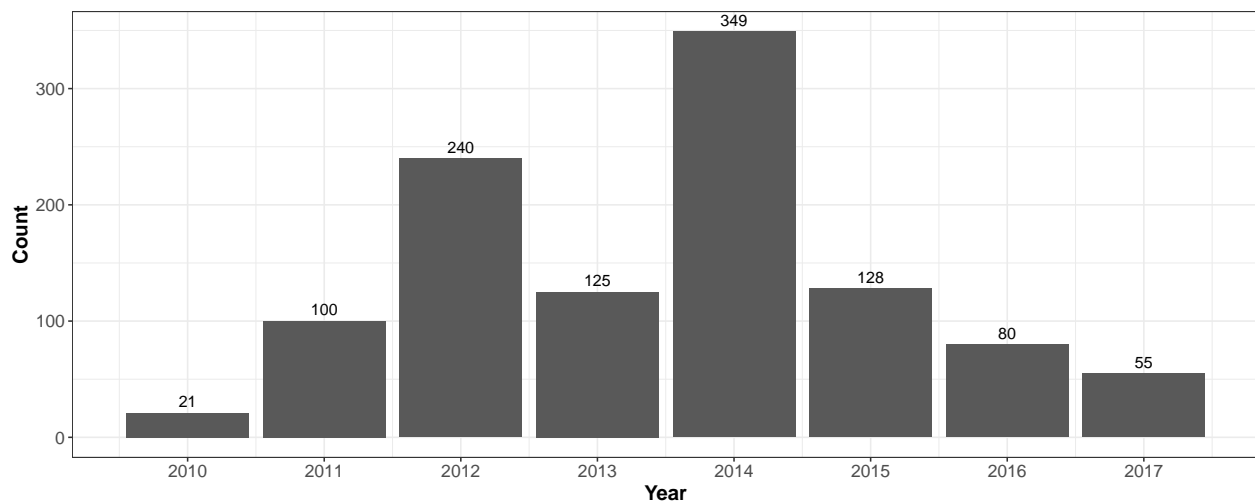
p



### Novel Sequence Types By Year

```
p <- ggplot(data=sequence_types, aes(x=year, y=novel)) +
  xlab("Year") +
  ylab("Count") +
  geom_bar(stat='identity') +
  geom_text(aes(label=novel), vjust = -0.5) +
  scale_x_continuous(breaks = round(seq(min(sequence_types$year), max(sequence_types$year), by = 1), 1)) +
  theme_bw() +
  theme(axis.text=element_text(size=12),
        axis.title=element_text(size=14,face="bold"))
```

p

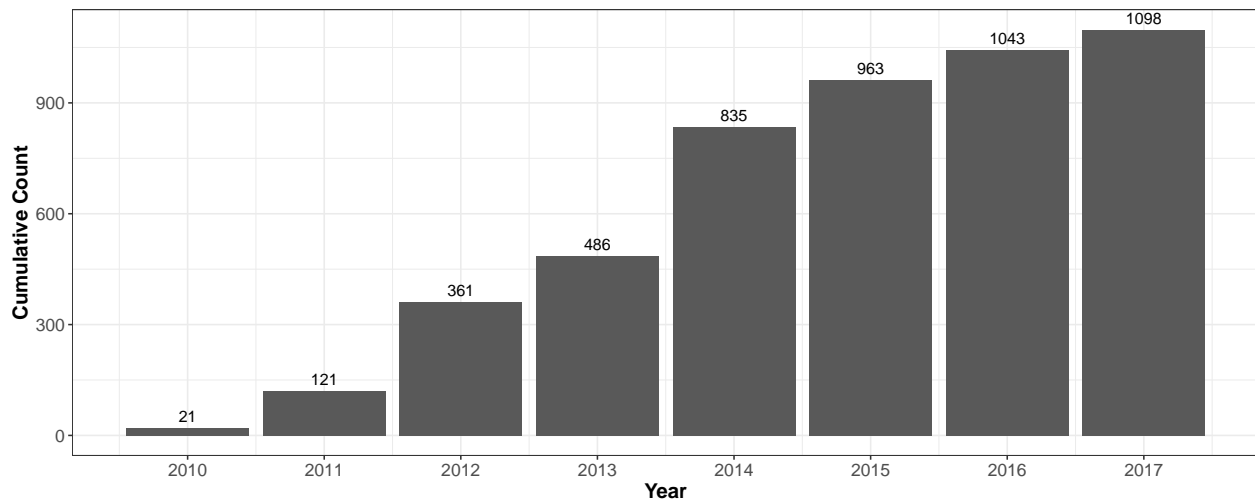


### Overall Novel Sequence Types By Year

```
p <- ggplot(data=sequence_types, aes(x=year, y=overall_novel)) +
  xlab("Year") +
  ylab("Cumulative Count") +
  geom_bar(stat='identity') +
  geom_text(aes(label=overall_novel), vjust = -0.5) +
  scale_x_continuous(breaks = round(seq(min(sequence_types$year), max(sequence_types$year), by = 1), 1))
```

```
theme_bw() +
theme(axis.text=element_text(size=12),
      axis.title=element_text(size=14,face="bold"))
```

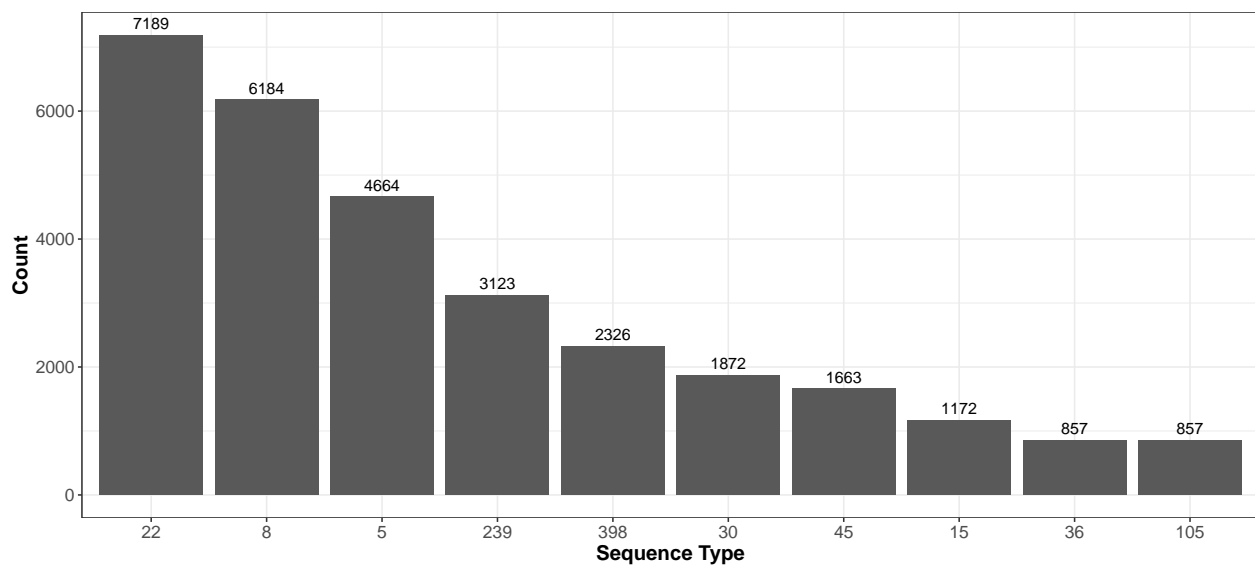
p



### Top 10 Sequence Types

```
p <- ggplot(data=top_st[1:10,], aes(x=reorder(st, -count), y=count)) +
  xlab("Sequence Type") +
  ylab("Count") +
  geom_bar(stat="identity") +
  geom_text(aes(label=count), vjust = -0.5) +
  theme_bw() +
  theme(axis.text=element_text(size=12),
        axis.title=element_text(size=14,face="bold"))
```

p



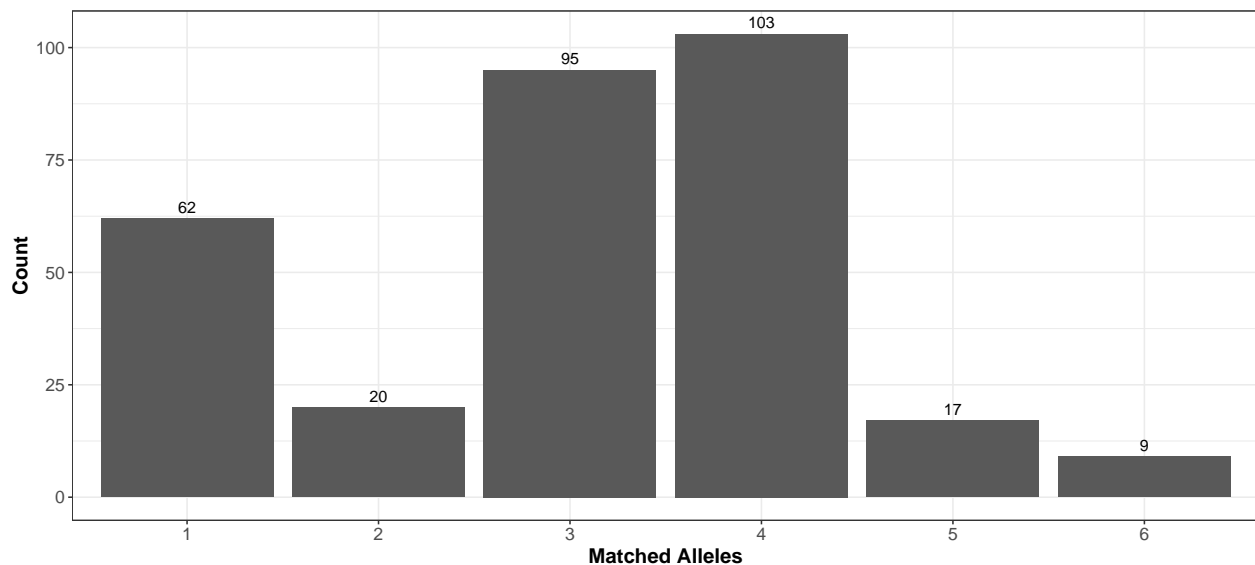
```
# Output plot to PDF and PNG
staphopia::write_plot(p, paste0(getwd(), '/images/figure-x-top-10-sequence-types'))
```

## Total Allele Matches For Unassigned Samples

```
allele_matches <- get_mlst_allele_matches(ps[ps$st == 0,]$sample_id)
df <- as.data.frame(table(allele_matches[allele_matches$matches < 7,]$matches))
colnames(df) <- c("matches", "count")
```

```
p <- ggplot(data=df, aes(x=matches, y=count)) +
  xlab("Matched Alleles") +
  ylab("Count") +
  geom_bar(stat="identity") +
  geom_text(aes(label=count), vjust = -0.5) +
  theme_bw() +
  theme(axis.text=element_text(size=12),
        axis.title=element_text(size=14,face="bold"))
```

p



## cgMLST Patterns

Finally, we'll look at cgMLST as a measure of genetic diversity. We will use the *get\_cgmlst()* function to get the cgMLST results for each Sample. This function might take a little while to retrieve all the results.

```
cgmlst <- get_public_cgmlst_patterns()
cgmlst$percent <- cgmlst$count / sum(cgmlst$total_samples)
cgmlst
```

##	samples_in_pattern	count	total_samples	percent
## 1	170	1	170	2.328343e-05
## 2	133	1	133	2.328343e-05
## 3	99	1	99	2.328343e-05
## 4	83	1	83	2.328343e-05
## 5	79	1	79	2.328343e-05
## 6	61	1	61	2.328343e-05
## 7	59	1	59	2.328343e-05
## 8	52	1	52	2.328343e-05
## 9	39	1	39	2.328343e-05
## 10	36	1	36	2.328343e-05

## 11	34	1	34 2.328343e-05
## 12	33	1	33 2.328343e-05
## 13	30	3	90 6.985029e-05
## 14	29	1	29 2.328343e-05
## 15	28	1	28 2.328343e-05
## 16	26	1	26 2.328343e-05
## 17	24	3	72 6.985029e-05
## 18	22	1	22 2.328343e-05
## 19	21	4	84 9.313372e-05
## 20	19	2	38 4.656686e-05
## 21	18	2	36 4.656686e-05
## 22	15	3	45 6.985029e-05
## 23	14	4	56 9.313372e-05
## 24	13	3	39 6.985029e-05
## 25	12	4	48 9.313372e-05
## 26	11	8	88 1.862674e-04
## 27	10	5	50 1.164171e-04
## 28	9	5	45 1.164171e-04
## 29	8	16	128 3.725349e-04
## 30	7	28	196 6.519360e-04
## 31	6	25	150 5.820857e-04
## 32	5	47	235 1.094321e-03
## 33	4	86	344 2.002375e-03
## 34	3	223	669 5.192205e-03
## 35	2	1363	2726 3.173531e-02
## 36	1	36827	36827 8.574588e-01

This gives us two columns:

1. `samples_in_pattern`: The number of samples with a given cgMLST pattern.
2. `count`: The number patterns with a given number of samples.
3. `total_samples`: Number of samples represented by a row (`samples_in_pattern * count`)
4. `percent`: Percent of samples represented

For example, if `samples_in_pattern` is 100 and the count is 2. That means there are **2** (count=2) cgMLST patterns that are shared by **100 samples** (`samples_in_count=100`) each, representing a total of **200 samples** (`count * samples_in_count`).

### Total Number of Distinct cgMLST Patterns

```
sum(cgmlst$count)
```

```
## [1] 38677
```

### How many shared cgMLST patterns?

```
sum(cgmlst[cgmlst$samples_in_pattern > 1, ]$count)
```

```
## [1] 1850
```

How many samples share a cgMLST pattern?

```
sum(cgmlst[cgmlst$samples_in_pattern > 1, ]$total_samples)
```

```
## [1] 6122
```

How many samples have a unique cgMLST pattern?

```
cgmlst$percent <- cgmlst$count / sum(cgmlst$total_samples)
cgmlst[cgmlst$samples_in_pattern == 1, ]
```

```
##      samples_in_pattern count total_samples  percent
## 36                      1 36827          36827 0.8574588
```

## Session Info

```
sessionInfo()
```

```
## R version 3.4.3 (2017-11-30)
## Platform: x86_64-pc-linux-gnu (64-bit)
## Running under: Ubuntu 16.04.2 LTS
##
## Matrix products: default
## BLAS: /usr/lib/libblas/libblas.so.3.6.0
## LAPACK: /usr/lib/lapack/liblapack.so.3.6.0
##
## locale:
##  [1] LC_CTYPE=en_US.UTF-8      LC_NUMERIC=C
##  [3] LC_TIME=en_US.UTF-8      LC_COLLATE=en_US.UTF-8
##  [5] LC_MONETARY=en_US.UTF-8  LC_MESSAGES=en_US.UTF-8
##  [7] LC_PAPER=en_US.UTF-8     LC_NAME=C
##  [9] LC_ADDRESS=C             LC_TELEPHONE=C
## [11] LC_MEASUREMENT=en_US.UTF-8 LC_IDENTIFICATION=C
##
## attached base packages:
## [1] stats      graphics  grDevices  utils      datasets  methods    base
##
## other attached packages:
## [1] scales_0.5.0  reshape2_1.4.3  ggplot2_2.2.1  staphopia_0.1.9
##
## loaded via a namespace (and not attached):
##  [1] Rcpp_0.12.15      knitr_1.20        magrittr_1.5
##  [4] munsell_0.4.3     colorspace_1.3-2  R6_2.2.2
##  [7] rlang_0.1.6       stringr_1.2.0     httr_1.3.1
## [10] plyr_1.8.4        tools_3.4.3       grid_3.4.3
## [13] data.table_1.10.4-3 gtable_0.2.0      htmltools_0.3.6
## [16] yaml_2.1.18       lazyeval_0.2.1    rprojroot_1.3-2
## [19] digest_0.6.15     tibble_1.4.2      curl_3.1
## [22] evaluate_0.10.1   rmarkdown_1.9     labeling_0.3
## [25] stringi_1.1.6     compiler_3.4.3    pillar_1.1.0
## [28] backports_1.1.2   jsonlite_1.5
```