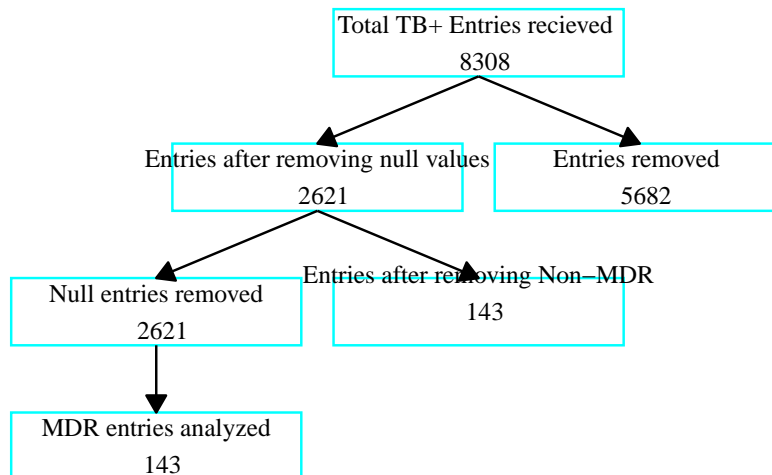


EPIDEMIOLOGY OF MULTI-DRUG
RESISTANT TB USING ISOLATES SENT TO
THE NATIONAL TUBERCULOSIS
REFERENCE LABORATORY (2015-2016)

METHODS

```
library('ggflowchart')
flow_chart <- tibble::tibble(from = c("Total TB+ Entries recieved
8308", "Total TB+ Entries recieved
8308", "Entries after removing null values
2621", "Entries after removing null values
2621", "Null entries removed
2621"),
                             to = c("Entries after removing null values
2621", "Entries removed
5682", "Entries after removing Non-MDR
143
", "Null entries removed
2621", "MDR entries analyzed
143"))
ggflowchart(flow_chart, colour = 'cyan', family = 'serif', x_nudge = 0.45)
```



R Packages used

```
#\ label: load-packages
suppressPackageStartupMessages(library('tidyverse'))

#install.packages('plotly')

#install.packages('ggmap')

#install.packages('forcats')
suppressPackageStartupMessages(library('ggmap'))
suppressPackageStartupMessages(library('plotly'))
suppressPackageStartupMessages(library('ggdist'))
suppressPackageStartupMessages(library('ggmap'))
```

FUNCTIONS

Fisher's function

```
Fishersfn <- function(df, var1, var2){  
  contingency_table <- table(df[[var1]], df[[var2]])  
  fisherExact <- fisher.test(contingency_table)  
  return (fisherExact)  
}
```

Getting percentages

```
pcts <- function(df, var_to_group_by) {  
  results <- df %>%  
    group_by_at(vars({{ var_to_group_by }})) %>%  
    summarise(counts = n()) %>%  
    mutate(Percentages = counts/sum(counts)*100)  
  return(results)  
}
```

Reading the file

```
mdr_tb_df <- read.csv('mdr_tbv4.csv')
```

Creating a new column to define MDR_TYPE

```
mdr_tb_df$MDR_TYPE <- NA
```

Here I have defined RpoB + katg as MDRKR (MDR KatG and RpoB), as well as those that are rpoB + katg + inhA

```
mdr_tb_dfv1 <- mdr_tb_df$MDR_TYPE[mdr_tb_df$RpoB == 'Resistant' & mdr_tb_df$KatG == 'Resistant']  
mdr_tb_df$MDR_TYPE[mdr_tb_df$RpoB == 'Resistant' & mdr_tb_df$KatG != 'Resistant' & mdr_tb_df$inhA == 'Resistant'] <- 'MDRKR'  
mdr_tb_dfv1 <- mdr_tb_df %>% select(-RpoB, -KatG, -inhA)
```

VISUALIZATIONS

Setting theme

```
theme_set(theme_gray())

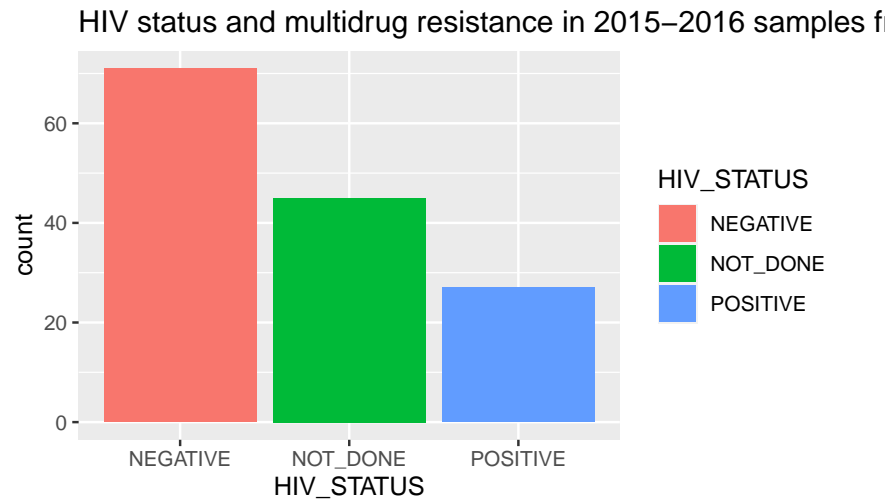
theme_update(

  plot.margin = margin(rep(20, 4))

)
```

HIV BAR PLOT

```
hivbar <- ggplot(mdr_tb_dfv1, mapping = aes(x=HIV_STATUS)) + geom_bar(mapping = aes(fill
hivbar
```



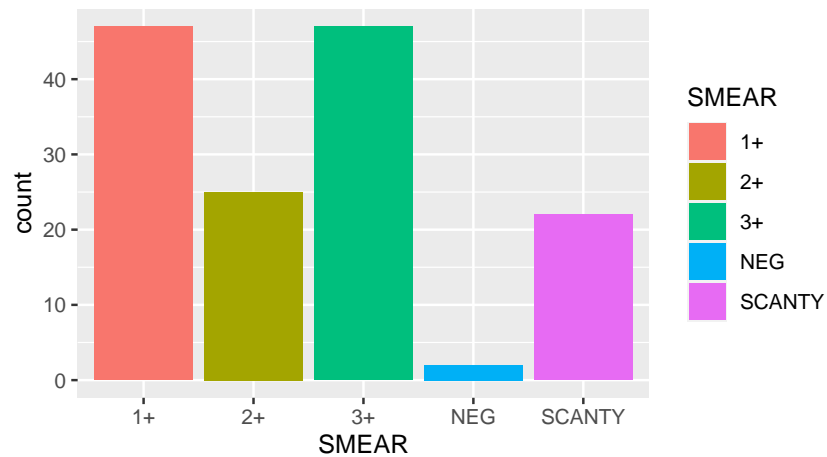
TB TYPE BAR PLOT

```
tb_type_bar <- ggplot(mdr_tb_dfv1, mapping = aes(x=TB_TYPE)) + geom_bar(mapping = aes(fill = S
```

SMEAR BAR PLOT

```
smearBar <- ggplot(mdr_tb_dfv1, mapping = aes(x=SMEAR)) + geom_bar(mapping = aes(fill = S  
smearBar
```

Smear results and multidrug resistance in 2015–2016 sample



Age groups based on WHO typical age divisions

```
mdr_tb_dfv3 <- mdr_tb_dfv1 %>% mutate(
  # Create categories
  age_group = dplyr::case_when(
    AGE <= 4 ~ "0-4",
    AGE > 4 & AGE <= 14 ~ "5-14",
    AGE > 14 & AGE <= 24 ~ "15-24",
    AGE > 24 & AGE <= 34 ~ "25-34",
    AGE > 34 & AGE <= 44 ~ "35-44",
    AGE > 44 & AGE <= 54 ~ "45-54",
    AGE > 54 & AGE <= 64 ~ "65+",
    AGE >= 64 ~ "35-44"
  ),
  # Convert to factor
  age_group = factor(
    age_group,
    level = c("0-4", "5-14", "15-24", "25-34", "35-44", "45-54", "65+")
  )
)
```


LOLLIPOP PLOT FOR COUNTY/GEOGRAPHICAL DISTRIBUTION

```
geoGrpsPcts <- pcts(mdr_tb_dfv1, 'Province')

GeoPops <- geoGrpsPcts %>%

  arrange(desc(Percentages)) %>%

  mutate( Province = forcats::fct_reorder(Province, Percentages)) %>%

  ggplot(aes(x=Province, y=Percentages)) +

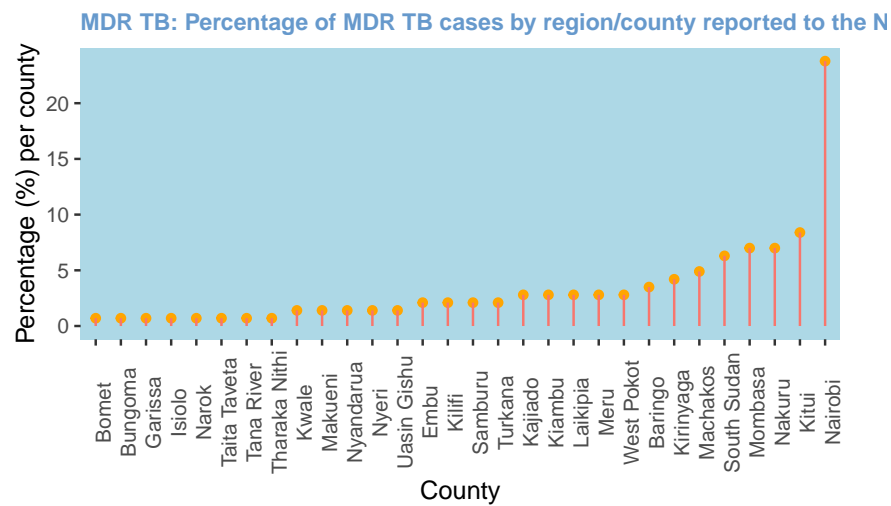
  geom_point(color = 'orange') +

  geom_segment(aes(x=Province, y=Percentages, xend=Province, yend=0, color = 'red')) +

  theme(axis.text.x = element_text(angle = 90), panel.background = element_rect(fill = 'lightblue'))

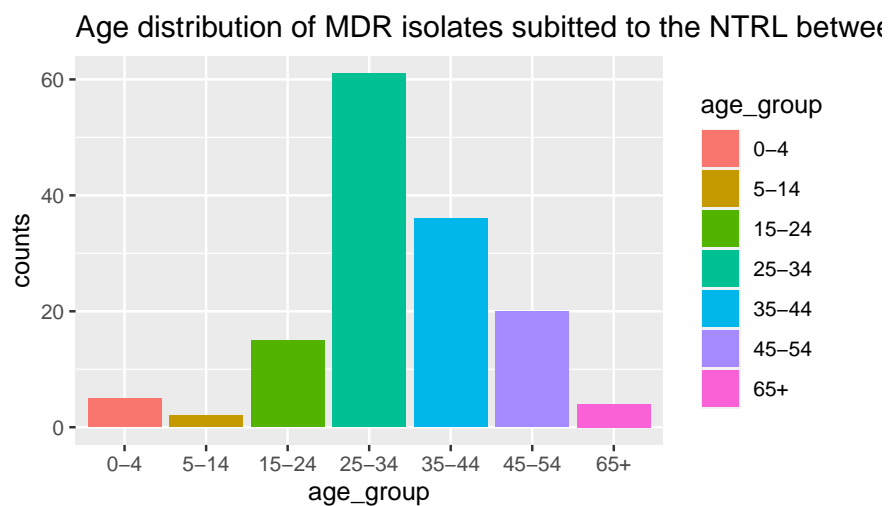
  labs(title = 'MDR TB: Percentage of MDR TB cases by region/county reported to the NTRL')

  guides(color = 'none')
GeoPops
```



AGE GROUP BAR PLOT

```
age_groupBars <- mdr_tb_dfv3 %>%  
  group_by(age_group) %>%  
  summarise(counts = n())  
  
ggplot() +  
  geom_bar(data = age_groupBars, aes(x = age_group, y = counts, fill = age_group), stat =  
  labs(title = 'Age distribution of MDR isolates subitted to the NTRL between 2015-2015')
```



```
age_groupBars  
  
# A tibble: 7 x 2  
  age_group counts  
  <fct>      <int>  
1 0-4         5  
2 5-14         2  
3 15-24        15  
4 25-34        61  
5 35-44        36
```

6	45-54	20
7	65+	4

INFERENCEAL STATISTICS

Fishers exact test for HIV status

```
p_value <- Fishersfn(mdr_tb_dfv3, 'HIV_STATUS', 'MDR_TYPE')  
p_value
```

Fisher's Exact Test for Count Data

```
data: contingency_table  
p-value = 0.2897  
alternative hypothesis: two.sided
```

Fisher's exact test for age

```
p_value <- Fishersfn(mdr_tb_dfv3, 'age_group', 'MDR_TYPE')  
p_value
```

Fisher's Exact Test for Count Data

```
data: contingency_table  
p-value = 0.175  
alternative hypothesis: two.sided
```

Fisher's exact test for COUNTY

```
p_value <- Fishersfn(mdr_tb_dfv1, 'Province', 'MDR_TYPE')  
p_value
```

Fisher's Exact Test for Count Data

```
data: contingency_table  
p-value = 0.06311  
alternative hypothesis: two.sided
```

Fisher's exact test for TB type

```
p_value <- Fishersfn(mdr_tb_dfv1, 'TB_TYPE', 'MDR_TYPE')  
p_value
```

Fisher's Exact Test for Count Data

```
data: contingency_table  
p-value = 0.9482  
alternative hypothesis: two.sided
```

Fisher's exact test for GENDER

```
p_value <- Fishersfn(mdr_tb_dfv1, 'GENDER', 'MDR_TYPE')  
p_value
```

Fisher's Exact Test for Count Data

```
data: contingency_table  
p-value = 0.3319  
alternative hypothesis: true odds ratio is not equal to 1  
95 percent confidence interval:  
 0.5015292 9.1443367  
sample estimates:  
odds ratio  
 2.088592
```

Fisher's exact test for SMEARS

```
p_value <- Fishersfn(mdr_tb_dfv1, 'SMEAR', 'MDR_TYPE')  
p_value
```

Fisher's Exact Test for Count Data

```
data: contingency_table  
p-value = 0.7377  
alternative hypothesis: two.sided
```

More characterizations

```
mdr_tb_dfv5 <- mdr_tb_df$MDR_TYPE[mdr_tb_df$RpoB == 'Resistant' & mdr_tb_df$KatG == 'Resi
```

Percentages of the MDR types

```
pcts(mdr_tb_df, 'MDR_TYPE')
```

```
# A tibble: 3 x 3
  MDR_TYPE counts Percentages
  <chr>      <int>      <dbl>
1 MDR+         7         4.90
2 MDRIR        11         7.69
3 MDRKR       125        87.4
```

TABLES OF DEMOGRAPHIC DISTRIBUTION OF MDR TYPES

```
table(mdr_tb_df$Province, mdr_tb_df$MDR_TYPE)
```

	MDR+	MDRIR	MDRKR
Baringo	1	0	4
Bomet	0	1	0
Bungoma	0	0	1
Embu	0	0	3
Garissa	1	0	0
Isiolo	0	0	1
Kajiado	1	0	3
Kiambu	0	0	4
Kilifi	0	0	3
Kirinyaga	0	1	5
Kitui	0	1	11
Kwale	0	0	2
Laikipia	0	0	4
Machakos	0	0	7
Makueni	0	0	2

Meru	1	1	2
Mombasa	0	0	10
Nairobi	2	1	31
Nakuru	0	0	10
Narok	0	0	1
Nyandarua	0	0	2
Nyeri	0	0	2
Samburu	0	1	2
South Sudan	1	2	6
Taita Taveta	0	0	1
Tana River	0	1	0
Tharaka Nithi	0	0	1
Turkana	0	0	3
Uasin Gishu	0	1	1
West Pokot	0	1	3

```
table(mdr_tb_df$HIV_STATUS, mdr_tb_df$MDR_TYPE)
```

	MDR+	MDRIR	MDRKR
NEGATIVE	2	3	66
NOT_DONE	3	5	37
POSITIVE	2	3	22

```
table(mdr_tb_df$SMEAR, mdr_tb_df$MDR_TYPE)
```

	MDR+	MDRIR	MDRKR
1+	2	5	40
2+	2	2	21
3+	1	2	44
NEG	1	0	1
SCANTY	1	2	19

```
table(mdr_tb_df$TB_TYPE, mdr_tb_df$MDR_TYPE)
```

	MDR+	MDRIR	MDRKR
FA_1STL_TR	1	3	31
FA_RE_TREA	0	0	3
MDR_CONTA	0	0	3
MDR_F	1	1	22
NEW	1	4	26
RE_AF_DEFA	0	1	13
SP_SM_NR	0	0	3

SP_SM_PR	4	2	23
UNKNOWN	0	0	1