Clinical, gene and mutation data of cancers with cryptic STMN2 events

STMN2 junction coverage

Overall STMN2 junction coverage in different cancer sites

```
## Warning in wilcox.test.default(c(64.8789804967288, 64.8789804967288,
## 51.8185835185835, : cannot compute exact p-value with ties
## Warning in wilcox.test.default(c(64.8789804967288, 64.8789804967288,
## 51.8185835185835, : cannot compute exact p-value with ties
```

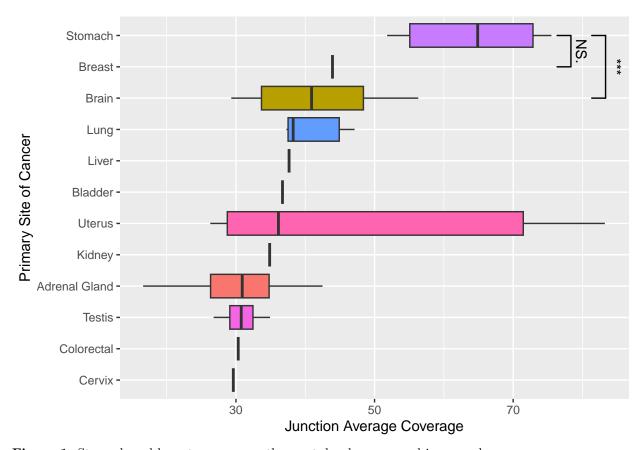


Figure 1: Stomach and breast cancers are the most deeply sequenced in general.

Cryptic STMN2 junction coverage in different cancer sites

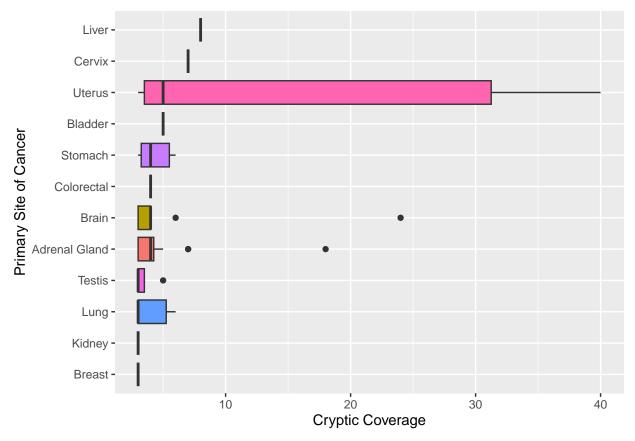


Figure 2:

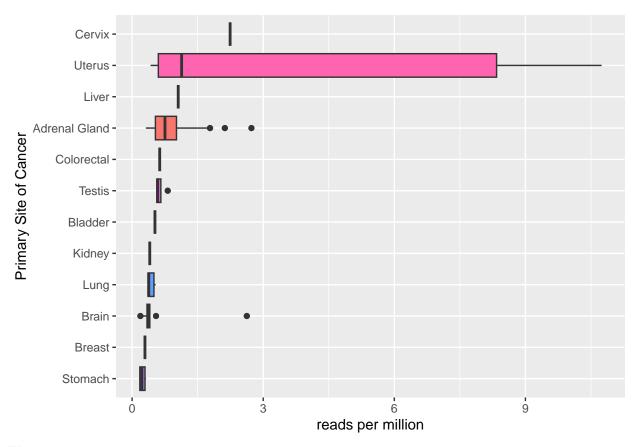


Figure 3:

On average, cancers in the stomach and breast have the greatest number of reads supporting cryptic STMN2 events. Cancers of the brain have significantly fewer reads supporting cryptic events.

Cancers of the cervix, uterus and liver have the greatest cryptic coverage. However, there is only one data value for cancers of the cervix and liver.

Cryptic STMN2 Expression

Cancers with STMN2 events (in general)



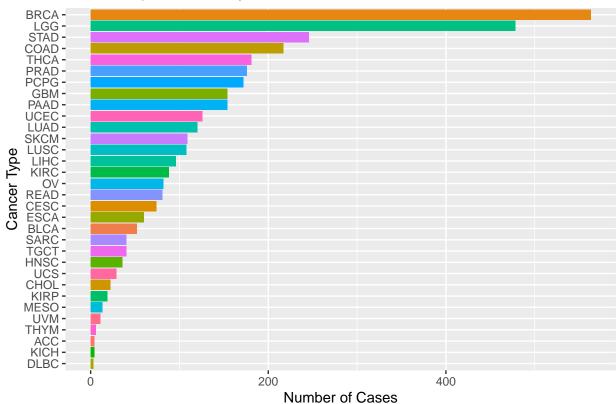
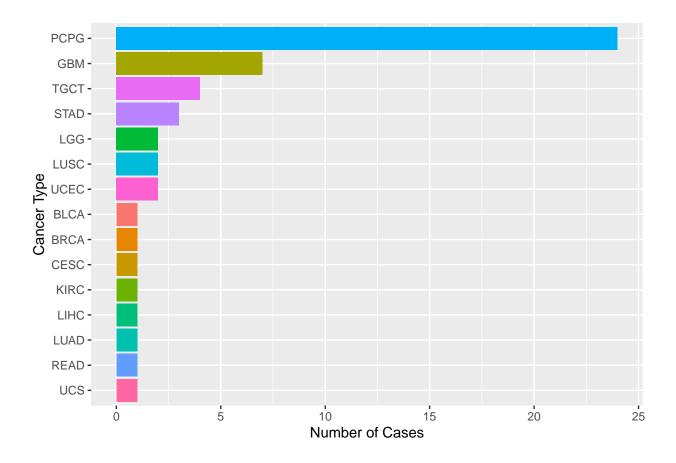
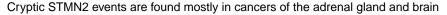


Figure 4: STMN2 events are found mostly in breast and brain cancer patients.



Primary sites of cancers

```
STMN2_clinical_jir_cryptic |>
  drop_na() |>
  ggplot(aes(x = fct_rev(fct_infreq(gdc_primary_site)))) +
  geom_bar(aes(fill = gdc_primary_site)) +
  coord_flip() +
  labs(
    x = "Primary Site of Cancer",
    y = "Number of Cases",
    title =
  "Cryptic STMN2 events are found mostly in cancers of the adrenal gland and brain"
  ) +
  theme(legend.position = "none", plot.title = element_text(size=9))
```



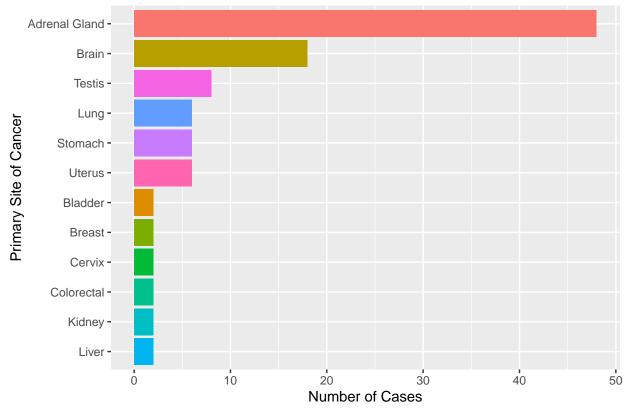


Figure 5: Cryptic STMN2 events are found mostly in the adrenal gland and brain.

Interestingly, Figure 5 shows that cryptic STMN2 expression is low in the breast. Comparing this to Figure 4, this shows that the STMN2 events expressed in breast cancers may be mostly annotated non-cryptic events.

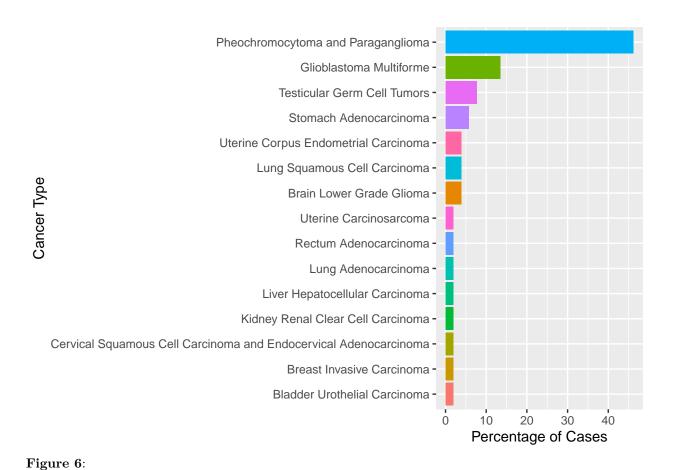
Which cancers have the most cryptic STMN2 events?

Fraction of each cancer with cryptic STMN2 expression

Table 1: Breast and brain cancers have high cryptic STMN2 expression.

| cancer | n | percent |
|--|----|-----------|
| Pheochromocytoma and Paraganglioma | 48 | 0.4615385 |
| Glioblastoma Multiforme | 14 | 0.1346154 |
| Testicular Germ Cell Tumors | 8 | 0.0769231 |
| Stomach Adenocarcinoma | 6 | 0.0576923 |
| Brain Lower Grade Glioma | 4 | 0.0384615 |
| Lung Squamous Cell Carcinoma | 4 | 0.0384615 |
| Uterine Corpus Endometrial Carcinoma | 4 | 0.0384615 |
| Bladder Urothelial Carcinoma | 2 | 0.0192308 |
| Breast Invasive Carcinoma | 2 | 0.0192308 |
| Cervical Squamous Cell Carcinoma and Endocervical Adenocarcinoma | 2 | 0.0192308 |
| Kidney Renal Clear Cell Carcinoma | 2 | 0.0192308 |
| Liver Hepatocellular Carcinoma | 2 | 0.0192308 |

| cancer | n | percent |
|------------------------|---|-----------|
| Lung Adenocarcinoma | 2 | 0.0192308 |
| Rectum Adenocarcinoma | 2 | 0.0192308 |
| Uterine Carcinosarcoma | 2 | 0.0192308 |



Where are the cancers with cryptic STMN2 events located?

Fraction of cases with STMN2 events in cancer sites

Table 2: Cancers with cryptic STMN2 events are found primarily in the adrenal gland and brain

| $gdc_primary_site$ | n | percent |
|----------------------|------|-----------|
| Adrenal Gland | 48 | 0.4615385 |
| Brain | 18 | 0.1730769 |
| Testis | 8 | 0.0769231 |
| Lung | 6 | 0.0576923 |
| Stomach | 6 | 0.0576923 |
| Uterus | 6 | 0.0576923 |
| Bladder | 2 | 0.0192308 |
| Breast | 2 | 0.0192308 |
| Cervix | 2 | 0.0192308 |

| gdc_primary_site | n | percent |
|------------------|---|-----------|
| Colorectal | 2 | 0.0192308 |
| Kidney | 2 | 0.0192308 |
| Liver | 2 | 0.0192308 |

Cancers with cryptic STMN2 events are primarily in the adrenal gland and brain

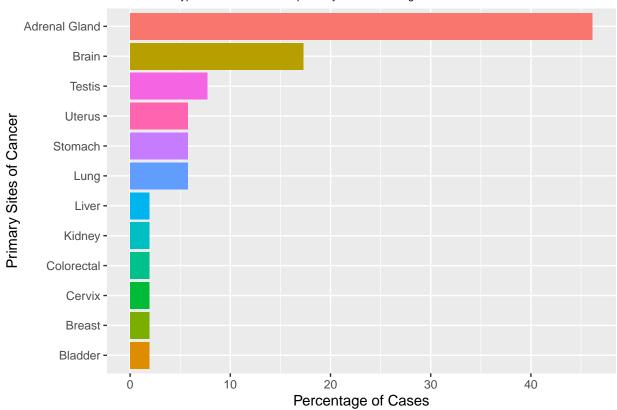


Figure 7:

TCGA/cBioPortal Clinical Data

Mutation Counts

UCEC and LUSC cancers have the greatest number of mutations

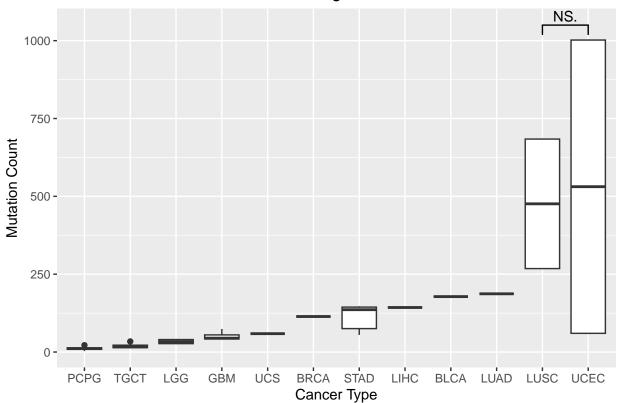


Figure 8: Among cancer patients with STMN2 cryptic expression, uterine corpus endometrial carcinoma (UCEC) has the greatest number of mutations.

Mutational burden

```
total_mutations_each_cancer <- cBio_clinical |>
    drop_na(Mutation.Count) |>
    filter(grepl("^\\d+$", Mutation.Count)) |>
    group_by(cancer_abbrev) |>
    summarise(total_mutations = sum(as.numeric(Mutation.Count)))

total_mutations_each_cancer_with_cryptic <- STMN2_cryptic_cBio |>
    drop_na(mutation_count) |>
    filter(mutation_count < 2500) |>
    group_by(cancer_abbrev) |>
    summarise(total_mutations_cryptic = sum(mutation_count))

mutations_each_cancer_general_vs_cryptic <- total_mutations_each_cancer |>
    left_join(total_mutations_each_cancer_with_cryptic, by=c("cancer_abbrev")) |>
```

Fraction of each cancer that has cryptic STMN2 events

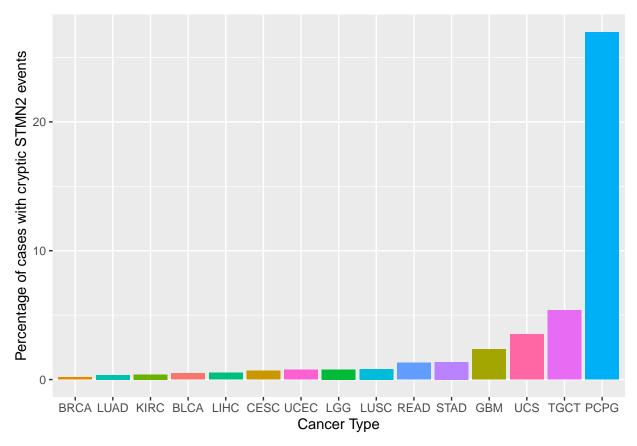
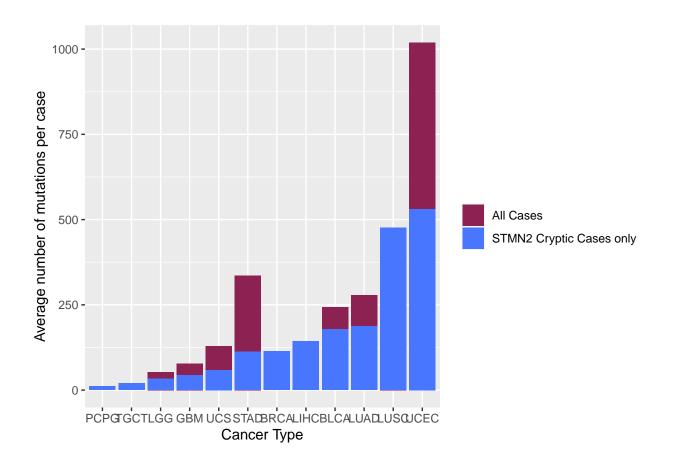


Figure 9:



```
## Warning in left_join(cBio_clinical, STMN2_clinical_jir, by = c("case_submitter_id")): Detected an un
## i Row 25 of 'x' matches multiple rows in 'y'.
## i Row 590 of 'y' matches multiple rows in 'x'.
## i If a many-to-many relationship is expected, set 'relationship =
## "many-to-many"' to silence this warning.

## Warning: Expected 1 pieces. Additional pieces discarded in 14659 rows [1, 2, 3, 4, 5, 6,
## 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, ...].

## Warning: There was 1 warning in 'mutate()'.
## i In argument: 'mutation_count = as.numeric(mutation_count)'.
## Caused by warning:
##! NAs introduced by coercion

## Warning: Computation failed in 'stat_signif()'
## Caused by error in 'if (scales$x$map(comp[1]) == data$group[1] | manual) ...':
##! missing value where TRUE/FALSE needed
```

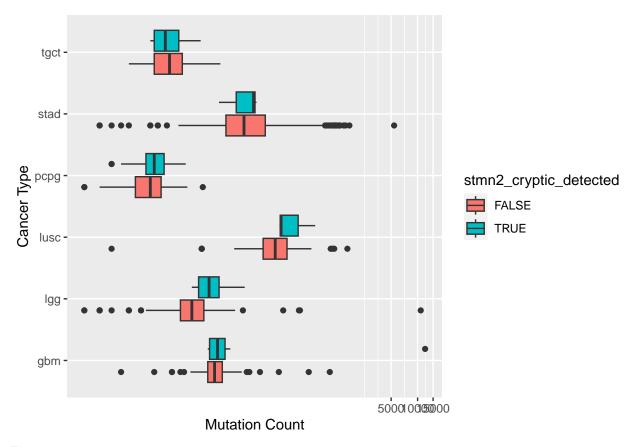


Figure 10:

Survival Comparisons

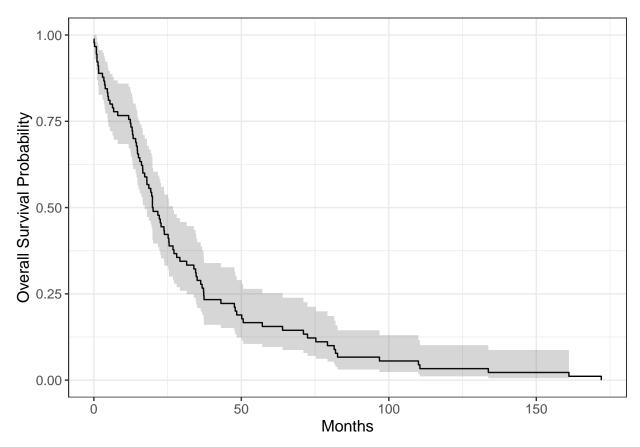


Figure 11:

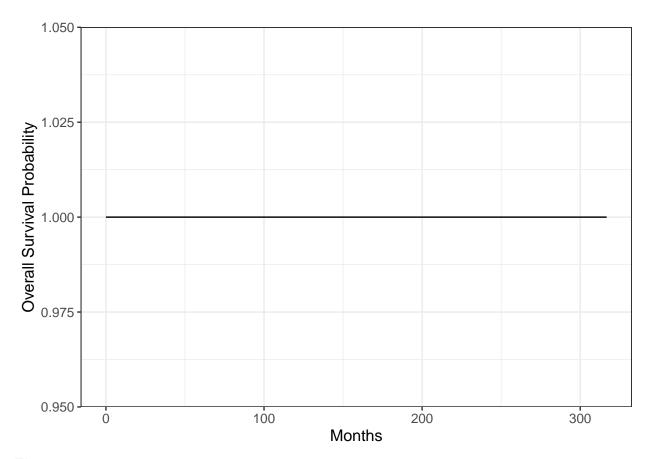


Figure 12:

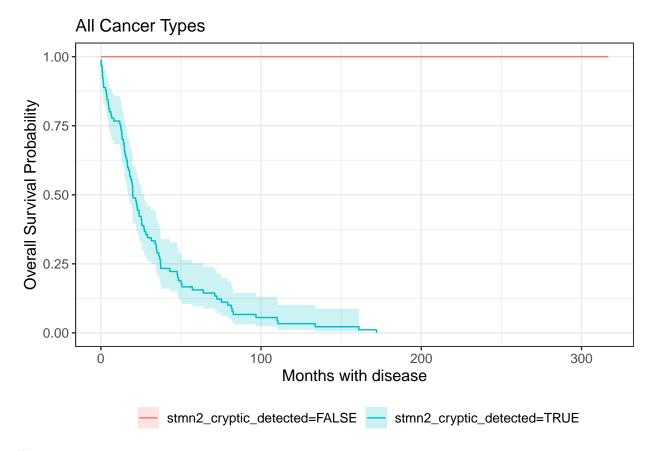


Figure 13:

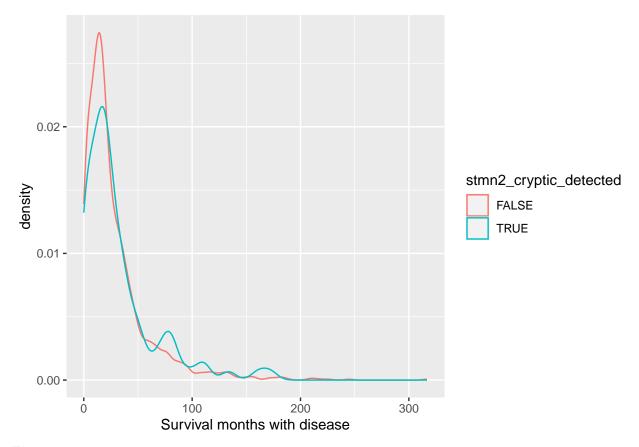


Figure 14:

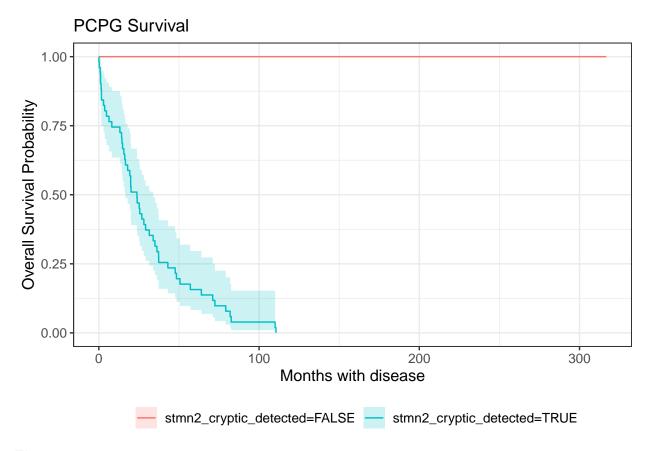


Figure 15:

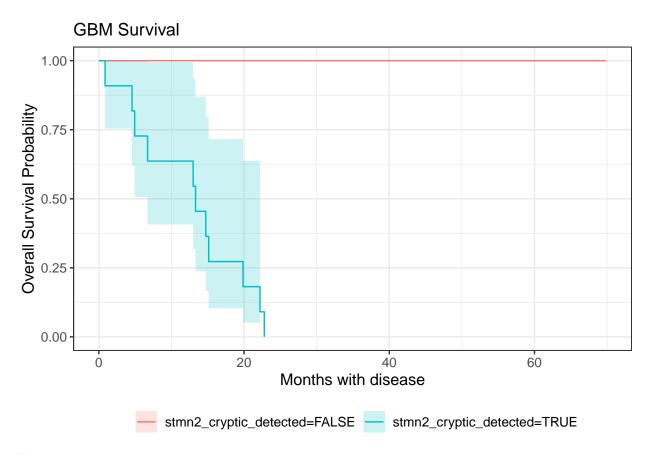


Figure 16:

Mutation Data of one patient

STMN2 cryptic expression

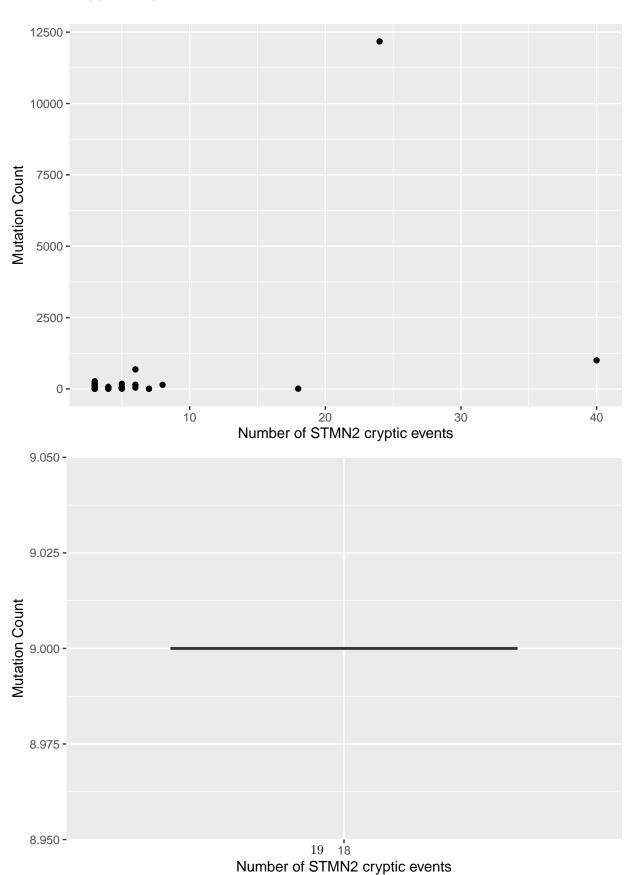


Figure 17: There is no correlation between number of STMN2 cryptic events and number of mutations.

Cancer Driver Genes

Mutations in each cancer gene

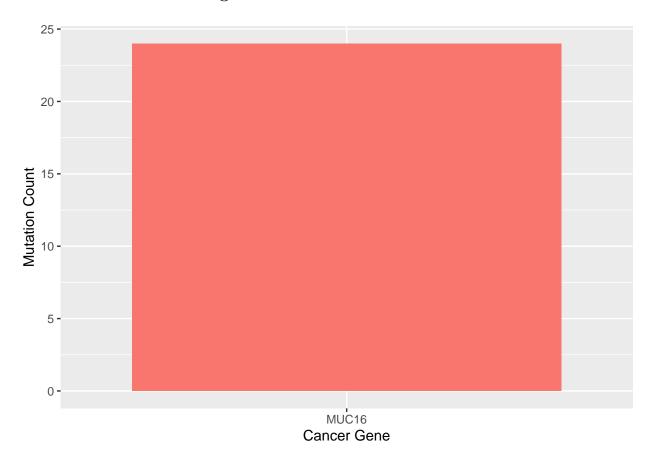


Figure 18: Ovarian cancer gene MUC16 has the greatest number of mutations.

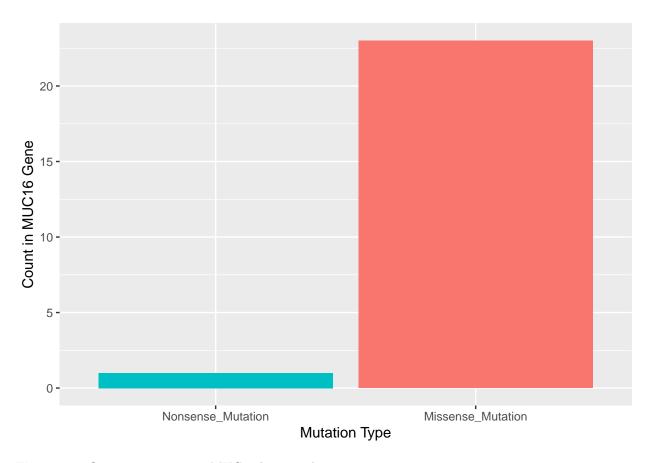


Figure 19: Ovarian cancer gene MUC16 has mostly missense mutations.

Types of cancer genes

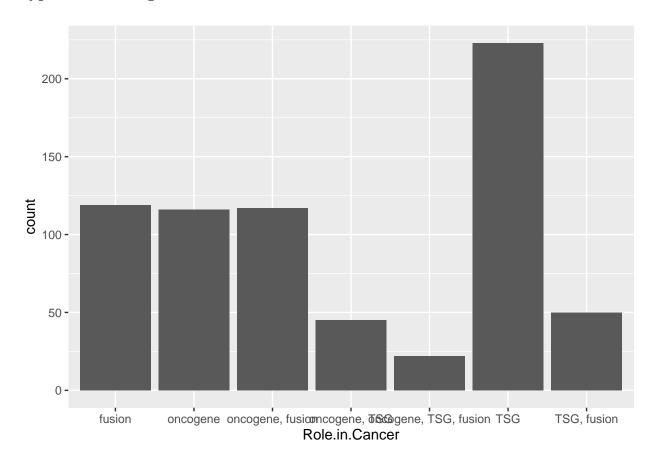


Table 3: 45% of cancer genes with cryptic STMN2 expression are tumour suppressor genes.

| TSG | n | percent |
|-----|-----|-----------|
| no | 394 | 0.5367847 |
| yes | 340 | 0.4632153 |

Table 4: 39% of cancer genes with cryptic STMN2 expression are oncogenes.

| oncogene | n | percent |
|----------|-----|-----------|
| no | 434 | 0.5912807 |
| yes | 300 | 0.4087193 |

Table 5: 44% of cancer genes with cryptic STMN2 expression are gene fusions.

| fusion | n | percent |
|--------|-----|-----------|
| no | 426 | 0.5803815 |

| fusion | n | percent |
|--------|-----|-----------|
| yes | 308 | 0.4196185 |

TCGA biolinks