Time to freezing of gait analysis: A time to event analysis

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Executive summary

In this project, we will be using data from kaggle. We aim at analyzing the expected duration of time until freezing of gait (FOG) occurs (time to FOG). FOG is a pattern occurring in patient with Parkinson disease. It is a debilitating symptom that afflicts many people with Parkinson's disease. Some events like start walking hesitation, turning body could be observed and help to detect FOG occurrence.

Introduction

We aim at analyzing the time to FOG in 44 Parkinson disease patients, with median age of 69 years old. Here FOG is the event of interest and it is expected within the execution of a given task in lab, or during daily life activity. We will analyse data from lab tasks. Our main goal is to analyse and explain overall FOG time to event model, the influence/importance of some features in the time to occurrence of FOG from the beginning to the end of the task. To achieve this, we model our data under kaplan-Meier model.

Data preparation, inspection and preprocessing.

let's join all metadata tables before diving into analysis.

- defog_metadata.csv Identifies each series in the tdcsfog dataset by a unique Subject, Visit, Test, Medication condition.
 - Visit Lab visits consist of a baseline assessment, two post-treatment assessments for different treatment stages, and one follow-up assessment.
 - Test Which of three test types was performed, with 3 the most challenging.
 - Medication Subjects may have been either off or on anti-parkinsonian medication during the recording.
- subjects.csv Metadata for each Subject in the study, including their Age and Sex as well as:
 - Visit Only available for subjects in the daily and defog datasets.
 - YearsSinceDx Years since Parkinson's diagnosis.
 - UPDRSIIIOn/UPDRSIIIOff Unified Parkinson's Disease Rating Scale score during on/off medication respectively. NFOGQ Self-report FoG questionnaire score. See: https://pubmed.ncbi.nlm.nih.gov/19660949/
- events.csv Metadata for each FoG event in all data series. The event times agree with the labels in the data series.
 - Id The data series the event occured in.
 - Init Time (s) the event began.
 - Completion Time (s) the event ended.
 - Type Whether StartHesitation, Turn, or Walking.
 - Kinetic Whether the event was kinetic (1) and involved movement, or akinetic (0) and static.
- tasks.csv Task metadata for series in the defog dataset. (Not relevant for the series in the tdcsfog or daily datasets.)
 - Id The data series where the task was measured.
 - Begin Time (s) the task began.
 - End Time (s) the task ended.

Task One of seven tasks types in the DeFOG protocol, described on this page.

Remove Visit from subjects table, as we do not use it

Note: we are going to consider only defog condition subjects.

Data structure

Ad event and tasks duration columns features:

```
    eventsDuration <- Completion - Init</li>
    tasksDuration <- Begin - end</li>
```

Data analysis

Subjects

How many unique subjects are in this dataset ?

There are 44 distinct subjects in this study.

Kinetic / Events

[1] 2232

#

How many trials has missing kinetic/status?

```
## [1] 0.48

48% of trials has missing Kinetic(status).
filter for missing kinetic entries/cases.

##
## new table dimension
```

19

```
## # A tibble: 3 x 19
##
     Ιd
                 Init Compl~1 Type Kinetic Begin
                                                    End Task Subject Visit Medic~2
##
     <chr>>
                <dbl>
                        <dbl> <chr>
                                      <dbl> <dbl> <chr> <chr>
                                                                       <dbl> <chr>
                                          1 1371. 1393. Turn~ ae2d35
## 1 02ea782681 1377.
                        1378. Turn
                                                                           2 on
## 2 02ea782681 1377.
                        1378. Turn
                                          1 1371. 1393. Turn~ ae2d35
                                                                           2 on
## 3 02ea782681 1466.
                        1467. Turn
                                          1 1461. 1472. Hots~ ae2d35
## # ... with 8 more variables: Age <dbl>, Sex <chr>, YearsSinceDx <dbl>,
       UPDRSIII_On <dbl>, UPDRSIII_Off <dbl>, NFOGQ <dbl>, eventsDuration <dbl>,
## #
       tasksDuration <dbl>, and abbreviated variable names 1: Completion,
```

What is the proportion of observed Kinetic events?

69 % of events has been observed.

2: Medication

Note: Given that each event is indicative of FOG, we will just gather them and consider that they form one class (Kinetic / events).

Age

What is the median age? Age numerical summary.

```
## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 28.00 61.00 69.00 67.44 73.00 82.00
```

At least half of subjects are 69 years old.

\mathbf{Sex}

Sex numerical summary

```
## Sex
## F M
## 0.45 0.55
```

There is almost 55.1498127% of men in this cohort.

Tasks

What kind of tasks has been performed?

```
## [1] "TUG-DT" "TUG-C" "Turning-DT" "Hotspot2" "Hotspot2-C" ## [6] "Hotspot1" "Turning-ST" "4MW" "4MW-C" "TUG-ST" ## [11] "Turning-C" "Hotspot1-C" "MB10" "MB11" "MB13" ## [16] "MB12"
```

how many tasks had been performed?

There had been 16 task performed.

Visit

How many rounds of visit did the patient had?

```
## [1] 2
```

What is the proportion of patient within each number of visit round?

```
## Visit
## 1 2
## 0.37 0.63
```

37% of patients had one Visit, while the remaining had 2.

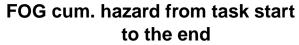
Medication

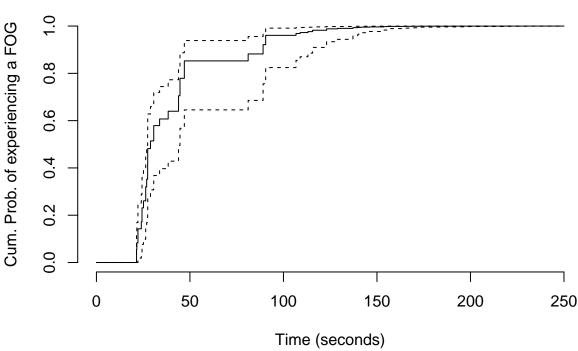
How many subjects are under parkinsonian-medication?

```
## Medication
## off on
## 68.89 31.11
```

31.11% are under medication.

Modelling and Analysis: overall time to FOG.





```
## Call: survfit(formula = Surv(Begin, End, Kinetic) ~ 1, data = fog)
##
## records n.max n.start events median 0.95LCL 0.95UCL
## [1,] 2232 190 36 1530 28.9 27.4 44.7
```

When considering time to the end of the task, half of patients experience FOG within 28.911 s.

Univariate analysis.

\mathbf{Sex}

p-value is 0.15 . There is no effect of sex on time to FOG.

Medication

P-value is 0 which is less than 0.05. There is huge effect of Medication in experiencing FOG holding other variables fixed.

Task type

```
## Warning in agreg.fit(X, Y, istrat, offset, init, control, weights = weights, :
## Loglik converged before variable 6; beta may be infinite.
## # A tibble: 3 x 3
##
                            estimate p.value
     term
##
     <chr>
                                <dbl>
                                       <dbl>
## 1 factor(Task)Hotspot1
                                1.03 0.0127
## 2 factor(Task)TUG-C
                                1.08 0.0186
                               -1.24 0.00165
## 3 factor(Task)Turning-DT
```

Three levels have significant effect on time to FOG, holding other variables fixed.

events Type

All p.values are greater than 0.05. There is no effect of Task type on FOG occurrence.

visit

p-value less than 0.05. Having more than one (1) round visit to hospital, has an influence on FOG occurrence.

yearSinceDx: year since diagnosis.

p-value is 0.5104841 which is > 0.05. There is no effect of Year since diagnosis on FOG occurrence.

Multivariate analysis

```
## # A tibble: 2 x 5
##
     term
                  estimate std.error statistic p.value
##
                      <dbl>
                                <dbl>
                                           <dbl>
                                                    <dbl>
     <chr>>
## 1 Medicationon
                      0.740
                               0.0702
                                           10.5 5.84e-26
## 2 Visit
                      0.240
                               0.0630
                                            3.81 1.36e- 4
```

Medication and Visit present a p value < 0.05. Medication increases the risk of FOG by 0.7 holding visit fixed, while, visit does the same with a rate of 0.2 while holding Medication fixed.

Results

This cohort has 44 subjects, half of the subjects are at least 69 years old. 31.11 % are under medication, 63% present more than 1 round visit to hospital. Our analysis shows that at least half of the subjects time to FOG is 28.911 seconds. Subjects under medication and having frequent visit to hospital are more likely to experience a FOG episode during a motor/kinetic task like gait.

Discussion

We aim at analyzing time to FOG in this study. Our analysis shows that FOG occurrence is related to Medication and Visit to hospital. Overall, half of subjects experience FOG within 28.9 seconds.

Conclusion

References

To be continued!!!!