## Notes on Longitudinal Data Analysis for Epidemiology

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## Welcome

## 1 State of the Art - Statistics

All this section is based on Twisk (2013)

### 1.1 Longitudinal studies

Longitudinal studies are defined as studies in which the outcome variable is repeateadly measured; i.e. the outcome variable is measured in the same subject on several occasions. – Extracted from Twisk (2013)

#### Characteristics:

- Observations of one subject over time are not independent of each other.
- Statistics should consider that repeated observations of each subject are correlated.
- This studies bring the illusion than are solving causality but only we can try temporality.

Table 1.1: Statistical notation

	Notation
number of subjects	i = 1 to N
number of covariates	j = 1  to  J
number of times a particular subject is measured	t = 1 to T
outcome variable	Y
covariates	X

#### 1.2 Cohort studies

#### 1.2.1 Observational cohort studies

The question of probable causality remains unanswered.

can be divided into:

<sup>\*</sup> prospective. \* The only one that can be characterized as longitudinal.

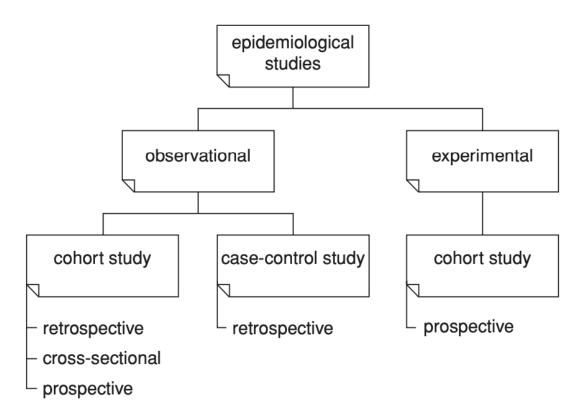


Figure 1.1: Image extracted from Twisk (2013)

- \* Analyze the longitudinal development of a certain characteristic over time (growth or deterioration).
- \* tracking: "stability" of a certain characteristic over time.
  - retrospective.
  - cross-sectional.

Term	Definition
age period	time from date of birth to date of measurement time or moment at which the measurement is taken
birth cohort	group of subjects born in the same year

#### 1.2.1.1 Confounding Effects

age, period and cohort effects could produce variations in the results.

- period effect.
  - If we measure physical activity during summers, it is likely that we can have more physical activity a hot summer than a rainy one. This can produce a bias in the age trend.
- **cohort effect**. If we want to unify results for the same age for cohorts that start at different ages, we will find that the trend is much flatter than the effects of the cohorts in isolation.

One way to avoid the bias is to use an approach called *multiple longitudinal design* Basically, multiple longitudinal design is to work with more than one cohort at the same time. If all the cohorts show a defined pattern for a particular measure in time, we will be able to detect it with this approach.

- Test or learning effect. Individuals start performing better with exposure.
- Low reproducibility of the measurements. Inter-period correlation coefficients (IPCs) (van 't Hofand Kowalski, 1979)

#### 1.2.2 Experimental cohort studies (clinical trials)

- There are prospective (ie longitudinal).
- The outcome variable Y is measured at least twice (the classical "pre-test," "post-test" design).
- The issue of causality can be covered

# 1.3 One continous outcome variable Y is repeatedly measured over time

#### 1.3.1 Two measurements

#### 1.3.1.1 Parametric: paired t-test

Is there a difference in the outcome variable Y between t = 1 and t = 2?

The **paired t-test** is used to test the hypothesis that the mean difference between Yt1 and Yt2 equals zero.

- Observations within one individual are dependent on each other.
- Use if the number of subjects is quite large (say above 25).

#### **1.3.1.1.1** Assumptions

- 1 The observations of different subjects are independent and;
- 2 The differences between the two measurements are approximately normally distributed.

#### 1.3.1.2 Non-parametric: (Wilcoxon) signed rank sum

Doesn't assume any distribution.

#### 1.3.2 More than two measurements

Does the outcome variable Y change over time?

#### 1.3.3 Multivariate analysis of variance (MANOVA) for repeated measurements

1 - Observations of different subjects. at each of the repeated measurements need to be independent; and. 2 - The observations need to be multivariate normally distributed, which is comparable but slightly more restrictive than the requirement that the differences between subsequent measurements be normally distributed.

#### 1.3.3.1 ANOVA (univariate) vs. MANOVA (multivariate)

To perform an ANOVA there is one extra assumption with 2 parts:

#### Sphericity assumption (epsilon coefficient)

- 3 All correlations in outcome variable Y between repeated measurements are equal, irrespective of the time interval between the measurements.
- 4 The variances of outcome variable Y are the same at each of the repeated measurements.

Which approach should be used? If the assumptions are met, ANOVA is more powerful for smaller samples:

"The restriction of the assumption of sphericity (i.e. equal correlations and equal variances over time) leads to an increase in degrees of freedom, i.e. an increase in power for the "univariate" approach. This increase in power becomes more important when the sample size becomes smaller. Historically, the "multivariate" approach was developed later than the "univariate" approach, especially for situations when the assumption of sphericity does not hold. So, one could argue that when the assumption of sphericity is violated, the "multivariate" approach should be used.

- Extracted from Twisk (2013)

MAMOVA in R - https://www.appsilon.com/post/manova-in-r

# 1.4 One continuous outcome variable Y is compared between different groups.

This design is known as the "one-within, one-between" design. Time is the within-subject factor and the group variable is the between-subjects factor.

Is there a difference in change over time for outcome variable Y between two or more groups?

- This question can also be answered with MANOVA for repeated measurements if it is assumed that the covariance matrices of the different groups that are compared to each other are homogeneous (independent sample t-test).
- Apparently, MANOVA could be biased when you have a lot of drop-offs in the study (Everitt (1998)).

#### 1.5 Continuous outcome variable and several covariates

(which can be either continuous, dichotomous, or categorical)

"traditional" methods tried to reduce the statistical longitudinal problem into a cross-sectional problem.

#### 1.5.1 Traditional methods.

- Analysis of the relationships between changes in different parameters between two points in time. (but, you are not using all the data)
- Use individual regression lines with time.

#### 1.5.2 New methods.

With the development of (new) statistical techniques, such as:

- 1. Generalized estimating equations (GEE) analysis and;
- 2. Mixed-model analysis,

it has become possible to analyze longitudinal relationships using all available longitudinal data, without summarizing the longitudinal development of each subject into one value.

Crippa (2022) A review of Longitudinal Data Analysis in R: https://rpubs.com/alecri/review\_longitudinal

- In this model the coefficients of interest are 1j, because these regression coefficients show the magnitude of the longitudinal relationship between the outcome variable (Yit) and the covariates (Xijt).
- Because of the dependency of the repeated observations within one subject, the relationship between X and Y must be adjusted for the subject. There is one by subject and are represented by dummy variables.

#### 1.5.2.1 GEE

- Before carrying out a GEE analysis, the within-subject correlation structure must be chosen.
- In the literature it is assumed that GEE analysis is robust against a wrong choice of the correlation structure. However, this is only the case when there are no missing data and when the model correctly specifies the mean.
- The interpretation of 1 is more complex than in a cross-sectional analysis.

#### 1.5.2.2 Mixed Models

- This type of study design is characterized by a hierarchical structure. In longitudinal studies observations within one subject over time are correlated. The observations over time are nested within the subject.
- The way mixed model analysis adjusts for the subject is different from the way GEE analysis adjusts for the subject.
- when an adjustment is made for the subject (i.e. the id\_number), for each subject different intercepts are calculated.
- The first step within a mixed model analysis is therefore to draw a normal distribution around the intercepts and in the second step the variance of that normal distribution is estimated.
- That variance is added to the longitudinal regression model in order to adjust for the subject in an efficient way.
- Because this variance is known as the **random intercept** (0i), mixed model analysis is also known as random coefficient analysis.
- The general idea behind a mixed model analysis is that the unexplained variance in outcome variable Y is divided into different components. One of the components is related to the random intercept and another component is related to random slopes.
- A model with a random intercept allows the intercepts to differ between the subjects, but the regression coefficient for the covariate X is the same for all the subjects. In a longitudinal study it is not uncommon that besides the intercepts also the regression coefficients for X differ between the subjects
- When regression coefficients for X differ between subjects, there is an interaction between the covariate X and the subject. As for the adjustment for the subject, also the interaction with the subject has to be added to a cross-sectional regression model with dummy variables.

#### **1.6 GAMM**

gamm considers that the observations aren't independent (not the same as gam)

#### 1.6.1 Random Effects

As we saw in the section about changing the basis, bs specifies the type of underlying base function. For random intercepts and linear random slopes we use bs = "re", but for random smooths we use bs = "fs".

3 different types of random effects in GAMMs (fac factor coding for the random effect; x0 continuous fixed effect):

- random intercepts adjust the height of other model terms with a constant value: s(fac, bs = "re")
- random slopes adjust the slope of the trend of a numeric predictor: s(fac, x0, bs = "re")
- random smooths adjust the trend of a numeric predictor in a nonlinear way: s(x0, fac, bs = "fs", m = 1), where the argument m=1 sets a heavier penalty for the smooth moving away from 0, causing shrinkage to the mean.

https://stats.stackexchange.com/questions/391912/how-to-fit-a-longitudinal-gam-mixed-model-gamm

- We can try to build a more appropriate model by fitting the data with a smoothed (non-linear) term specified by expressions of the form s(x)
- Radj is the variance explained by the model

## 2 Scientific articles

stringhini, 2018

Premature mortality reduction from chronic diseases

Biological Risk factors.

- high blood pressure.
- obesity.
- tobacco use.
- excess salt intake.
- diabetes.
- insufficient physical activity.
- alcohol consumption.

Socioeconomic status.

- occupational group.
- educational attainment.
- level of income and wealth.
- place of residence.

## 2.1 General Additive Mixed Model (GAMM)

#### semi-parametric model

Let's start with an equation for a Gaussian linear model:

$$y = 0 + x1 + 1 + N(0, 2)$$

What changes in a GAM is the presence of a smoothing term:

$$y = 0 + f(x1) + N(0, 2)$$

This term could be many things.

#### 2.1.1 Walking speed and age

Fixed Effects Predictors.

- age.
- height.

Random Effect.

• study at the intercept and age slope.

#### 2.1.2 Number of years of functioning lost (primary outcome)

It is based on the predictions of the previous model.

Fixed Effects

- age.
- age2.
- height.
- year of birth.
- distances walked.
- risk factor under study (minimally adjusted model).
- all risk factors (mutually adjusted models).

CI - Model based parametric 5000 bootstrap samples

#### 2.1.3 Years of life lost (secundary outcome)

Difference between the areas of the survival curves.

Survival curves Kaplan-Meier adjusted curves, conditional on survival to age 60 years. They run a shared frality Cox model with age as time scale, stratified by the levels of the given risk factor and a year of birth as covariate (for minimally adjusted models) or year of birth and the remaining risk factors as covariates (mutually adjusted models)

schrempft 2022

### 2.2 Pace of aging

similar to Dunedin Study investigators

- 1 Biomarkers were standarized for healthy men and women. Z-scores were reversed for HDL and creatinine clearence.
- 2 Mixed-effects models with a random interecept and a random linear slope, were used to calculate participants' personal slopes (change in biomarkers per year) Each year included time, age at baseline center in the samples mean, and a interaction term between the time and age at the baseline. For biomarkers that show a non linear trajectory, an additionally
- 3 The individual slopes for each biomarker (annual change in biomarker Z-score) wee aggregated to create a total Pace of Aging score.

Covariates: - Alcohol > 14 units. - hipertensive or diabetic medication. - pysical inactivity. - smoking status

carmeli, 2019

## 3 Data Analysis

```
library(haven) # read stata file format
library(readx1) # read excel file format
library(kableExtra)
library(lubridate)
library(tidyr)
library(plotly)
library(ggplot2)
library(ggbeeswarm)
library(PerformanceAnalytics)
library(forcats)
library(gamm4)
library(purrr)
```

The data was extracted from the Canadian Longitudinal Study on Aging Webpage.

The baseline study was conducted in 2008, the first follow-up in 2015. It seems to be data for the second follow up (variables ending in \_COF2), even if there is not information available in the webpage.

#### 3.1 Variables

I will assume that for the second follow-up, the variable names are the same as those described for the first follow-up.

Some of the variables are labeled. I will extract the labels to can use the information later. For this I create dlabel().

```
# Extract labels and save them in small datasets
dlabel <- function(variable){
data |>
    dplyr::select({{variable}}) |>
    dplyr::mutate(lab = haven::as_factor({{variable}})) |>
    dplyr::filter (!duplicated({{variable}})) |>
    dplyr::arrange({{variable}})
}

ed_label <- dlabel(ED_UDR04_COM)
sdc_label <- dlabel(SDC_MRTL_COM)
inc_tot_label <- dlabel(INC_TOT_COM)
inc_ptot_label <- dlabel(INC_PTOT_COM)
wea_label <- dlabel(WEA_SVNGSVL_MCQ)</pre>
```

This are the variables available in the dataset:

```
table <- rbind(var_baseline, var_follow_up1) |>
  dplyr::select(name, label, everything(), -unit, -table) |>
  dplyr::arrange(name) |>
  kableExtra::kable(caption = "Variables in the dataset") |>
  kableExtra::kable_classic(full_width = F)
table
```

Table 3.1: Variables in the dataset

label	valueType	question
Participant age at beginning of FU1	integer	NA
Age (years)	integer	NA
Highest Level of Education - Respondent, 4 Levels	text	NA
Total personal income	text	What is your
Total household income	text	What is you
Marital/partner status	text	What is you
Sex	text	Are you male
Total value of savings and investments	text	What is the
Total value of savings and investments	text	What is the
Total time required to complete 4mWalk (in seconds)	decimal	NA
Total time required to complete 4mWalk (in seconds)	decimal	Record time
In-Home Questionnaire start date& time	datetime	NA
Date and time at start of interview	datetime	NA
	Participant age at beginning of FU1 Age (years) Highest Level of Education - Respondent, 4 Levels Total personal income Total household income Marital/partner status Sex Total value of savings and investments Total value of savings and investments Total time required to complete 4mWalk (in seconds) Total time required to complete 4mWalk (in seconds) In-Home Questionnaire start date& time	Participant age at beginning of FU1 integer  Age (years) integer  Highest Level of Education - Respondent, 4 Levels text  Total personal income text  Marital/partner status text  Sex text  Total value of savings and investments text  Total value of savings and investments text  Total time required to complete 4mWalk (in seconds) decimal  Total time Questionnaire start date& time datetime

### 3.2 Exploratory Data Analysis (EDA)

#### 3.2.1 General observations

- In this dataset there are are reported the baseline values (\*\_COM), as well as the first follow up in 2017/18 (\*\_COF1) and the second follow up (\*\_COF2), that seems to be 2019/20/21 in the data.
- Data were collected between 2012 and 2021.
- There are 1000 observations in the dataset. In the baseline study 47.9 % are men and 52.1 % are women.

#### summary(data)

Min.

:45.00

Min.

:47.00

id	startdate_COM	startdate_COF1	startdate_COF2
Min. : 6	Length: 1000	Length:1000	Length:1000
1st Qu.:21704	Class :character	Class :character	Class :character
Median :45524	Mode :character	Mode :character	Mode :character
Mean :44858			
3rd Qu.:67622			
Max. :90271			
AGE_NMBR_COM	AGE_NMBR_COF1	AGE_NMBR_COF2 SEX_A	ASK_COM

Min.

:50.00

Length: 1000

```
Class : character
1st Qu.:55.00
                1st Qu.:58.00
                                 1st Qu.:60.00
Median :63.00
                Median :65.00
                                 Median :68.00
                                                  Mode :character
       :63.47
                        :66.06
Mean
                Mean
                                 Mean
                                         :68.44
3rd Qu.:72.00
                3rd Qu.:74.00
                                 3rd Qu.:76.00
Max.
       :86.00
                Max.
                        :89.00
                                 Max.
                                         :91.00
                NA's
                        :81
                                 NA's
                                         :148
ED UDRO4 COM
                 SDC_MRTL_COM
                                  INC TOT COM
                                                   INC PTOT COM
Min.
       :1.000
                Min.
                        :1.000
                                 Min.
                                         :1.000
                                                  Min.
                                                          :1.000
1st Qu.:4.000
                1st Qu.:2.000
                                 1st Qu.:2.000
                                                  1st Qu.:2.000
Median :4.000
                Median :2.000
                                 Median :3.000
                                                  Median :2.000
Mean
       :3.577
                        :2.308
                                         :3.494
                                                          :2.792
                Mean
                                 Mean
                                                  Mean
3rd Qu.:4.000
                 3rd Qu.:2.000
                                 3rd Qu.:4.000
                                                  3rd Qu.:3.000
       :9.000
                        :5.000
                                         :9.000
                                                          :9.000
Max.
                Max.
                                 Max.
                                                  Max.
WEA_SVNGSVL_MCQ
                 WLK_TIME_COM
                                    WLK_TIME_COF1
                                                         WLK_TIME_COF2
                                           :-99999.00
Min.
       :1.000
                        :-88.000
                                   Min.
                                                                :-99999.00
                Min.
                                                         Min.
1st Qu.:2.000
                1st Qu.: 3.600
                                   1st Qu.:
                                                 3.60
                                                         1st Qu.:-99993.00
Median :3.000
                Median: 4.090
                                                 4.09
                                   Median:
                                                         Median:
                                                                      3.22
Mean
       :3.116
                        : 4.172
                                           : -6439.67
                                                                :-46395.00
                Mean
                                   Mean
                                                         Mean
3rd Qu.:3.000
                3rd Qu.: 4.683
                                    3rd Qu.:
                                                 4.72
                                                         3rd Qu.:
                                                                      4.18
Max.
       :9.000
                Max.
                        : 14.410
                                   Max.
                                                10.63
                                                         Max.
                                                                     20.32
NA's
       :66
                NA's
                        :4
                                    NA's
                                           :81
                                                         NA's
                                                                :148
```

- The cases were NA values were reported were for COF1 (0%) and COF2 (0%) of the original database. These rows were removed. The NAs for WEA\_SVNGSVL\_MCQ were also eliminated from the analysis.
- Variable height, that can correlate with walking speed is missing.

```
is.na(data$startdate_COF1) == FALSE,
is.na(data$startdate_COF2) == FALSE,
is.na(data$WEA_SVNGSVL_MCQ) == FALSE) # remove NAs
```

#### 3.2.2 Data wrangling

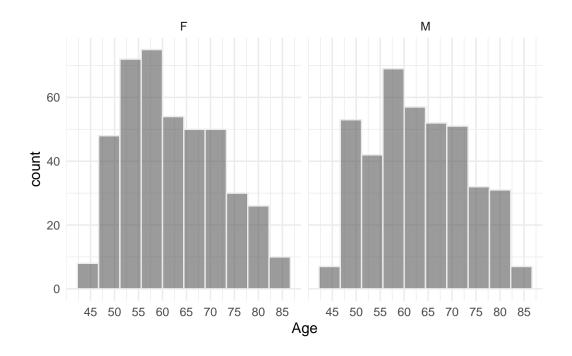
- The dataset is splitted in subtables to make it easier to use.
- The variable Walking time is converted to Walking speed.
- Following Stringhini et al. (2018), the ages have been reduced from 45 to 90.

```
# Socieconomic variables sub-table
soc vars <- data |>
 haven::zap_labels() |> # I have already saved the labels in tables
  tidyr::pivot longer(col = c(9, 11, 12, 13),
                      values_to = "SOC_VALS",
                      names_to = "SOC_VARS") |>
  dplyr::filter(!SOC_VALS < 0) |>
  dplyr::select(id, SOC_VALS, SOC_VARS)
# Gender sub-table
gender <- data |>
 haven::zap_labels() |>
  dplyr::select(id, SEX_ASK_COM)
# Walking speed sub-table
wlk_time <- data |>
 haven::zap_labels() |>
  tidyr::pivot_longer(14:16, values_to = "WLK_TIME", names_to = "COHORT") |>
  dplyr::filter(!WLK TIME < 0) |>
  tidyr::separate(COHORT, into = c("A", "B", "COHORT"), sep = "_") |>
  dplyr::select(id, COHORT, WLK_TIME)
# Age sub-table
age <- data |>
 haven::zap_labels() |>
  pivot_longer(5:7,
               values_to = "AGE_VALS", names_to = "AGE_VARS") |>
  dplyr::select(id, AGE_VALS, AGE_VARS) |>
  dplyr::filter(!is.na(AGE_VALS)) |>
  tidyr::separate(AGE_VARS, into = c("A", "B", "COHORT"), sep = "_") |>
```

```
dplyr::select(id, COHORT, AGE_VALS)
  # Final table
  data_by_cohort <- dplyr::left_join(age, wlk_time)</pre>
Joining with `by = join_by(id, COHORT)`
  data_plot <- dplyr::left_join(data_by_cohort, soc_vars)</pre>
Joining with `by = join_by(id)`
Warning in dplyr::left_join(data_by_cohort, soc_vars): Detected an unexpected many-to-many re
i Row 1 of `x` matches multiple rows in `y`.
i Row 1 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.
  # Convert Walking time to Walking speed
  data_plot <- data_plot |>
    dplyr::mutate(WLK_TIME = (4 / WLK_TIME)) |>
    dplyr::filter(AGE_VALS >= 45,
                   AGE_VALS <= 90)
    data_plot |>
      dplyr::left_join(gender) |>
      select(id, AGE_VALS, SEX_ASK_COM) |>
      filter (!duplicated(id)) |>
      ggplot() +
      scale_fill_brewer() +
      geom_histogram(
        aes(AGE_VALS),
        bins = 10,
        color = "#e9ecef",
        alpha = 0.6,
        position = 'identity'
      scale_x_continuous(breaks = round(seq(45,90, by = 5),1)) +
      xlab("Age") +
```

```
facet_wrap(vars(SEX_ASK_COM)) +
theme_minimal()
```

Joining with `by = join\_by(id)`



#### Comments:

• There are less observations for older ages.

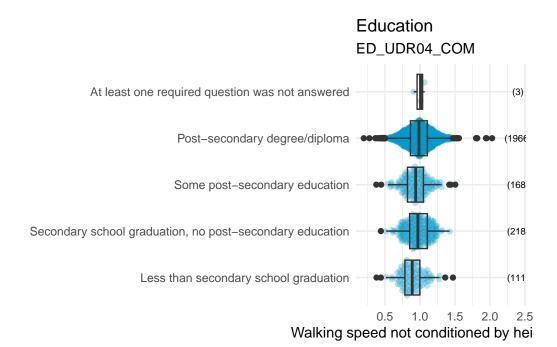
#### 3.2.3 Socioeconomic Variables

```
data_boxplot <- data_plot |>
   dplyr::filter(SOC_VARS == 'ED_UDR04_COM') |>
   dplyr::left_join(ed_label, by = dplyr::join_by(SOC_VALS == ED_UDR04_COM))
# Number of points by category
   obs1 <- data_boxplot |> dplyr::count(lab)

data_boxplot |>
   ggplot(aes(x = as.factor(lab), y = WLK_TIME)) +
   geom_quasirandom(alpha = 0.3, color = "deepskyblue3") +
```

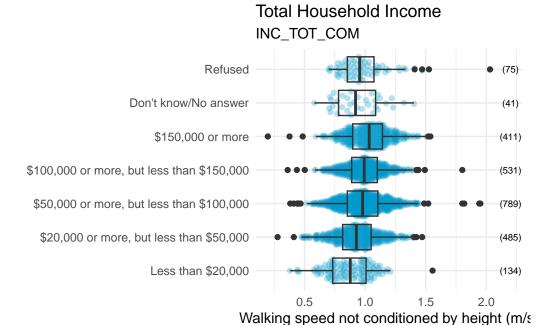
Warning: Removed 407 rows containing non-finite values (`stat\_boxplot()`).

Warning: Removed 407 rows containing missing values (`position\_quasirandom()`).



Warning: Removed 407 rows containing non-finite values (`stat\_boxplot()`).

Warning: Removed 407 rows containing missing values (`position\_quasirandom()`).



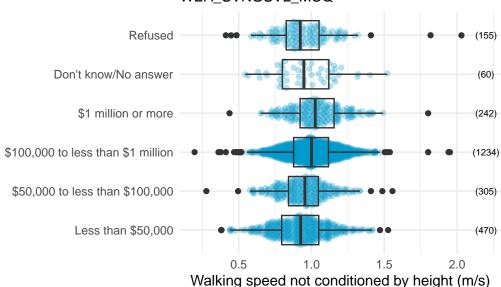
data\_boxplot <- data\_plot |>
 dplyr::filter(SOC\_VARS == 'WEA\_SVNGSVL\_MCQ') |>

```
dplyr::left_join(wea_label,
                   by = dplyr::join_by(SOC_VALS == WEA_SVNGSVL_MCQ))
# Number of points by category
obs3 <- data_boxplot |> dplyr::count(lab)
data_boxplot |>
 ggplot(aes(x = as.factor(lab), y = WLK_TIME)) +
 geom_quasirandom(alpha = 0.3, color = "deepskyblue3") +
 geom_boxplot(fill = 'transparent') +
 coord_flip() +
 labs(title = "Total Value of Savings and Investments",
       subtitle = "WEA_SVNGSVL_MCQ") +
 xlab("") +
 ylab("Walking speed not conditioned by height (m/s)") +
    geom_text(data = obs3, aes(y = 2.2,
                             label = paste0("(",n,")")),
                             size = 2.5) +
 theme_minimal()
```

Warning: Removed 407 rows containing non-finite values (`stat\_boxplot()`).

Warning: Removed 407 rows containing missing values (`position\_quasirandom()`).

# Total Value of Savings and Investments WEA\_SVNGSVL\_MCQ



Considering that we don't have the height values, we will make comments solely about the

Comments:

• 79.7% of the participants hold a post-secondary degree/diploma.

amount of observations in each category instead of analyzing the walking speed.

- 50of the participants have saving and investments between \$100,000 and \$1 million dollars
- However, the values of total income are more homogeneous among categories, being the maximum 32%. This could indicate that some study participants with a post-secondary degree and \$100,000 \$1 million dollars of savings and investments come from more than one total household income category. More analysis should be carry out to confirm this.
- Data reported as 'refused' or 'No answer' should be removed of the analysis.

#### 3.2.4 Walking Speed as function of age.

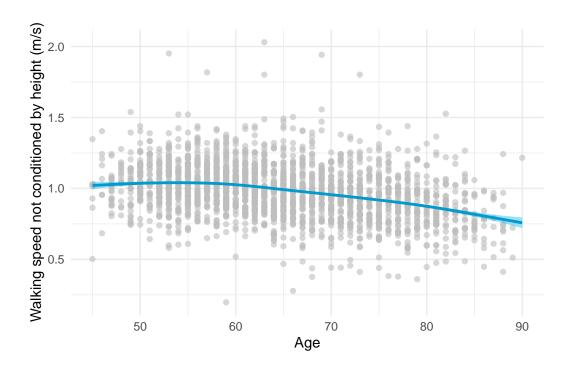
- Obervations are not independent
- Outcome variable: walking speed

First, let's visualize the data. ggplot2 function geom\_smooth() adjusts the data to walking-speed ~ s(age, bs = "cs").

'geom\_smooth()' using method = 'gam' and formula = 'y ~ s(x, bs = "cs")'

Warning: Removed 1628 rows containing non-finite values (`stat\_smooth()`).

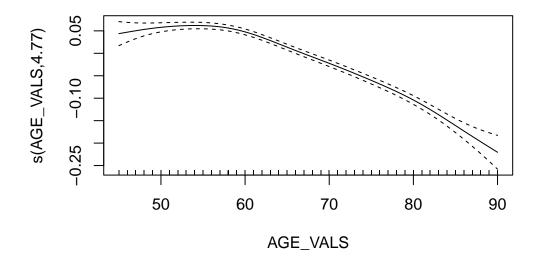
Warning: Removed 1628 rows containing missing values (`geom\_point()`).



If we fit this basic first model:

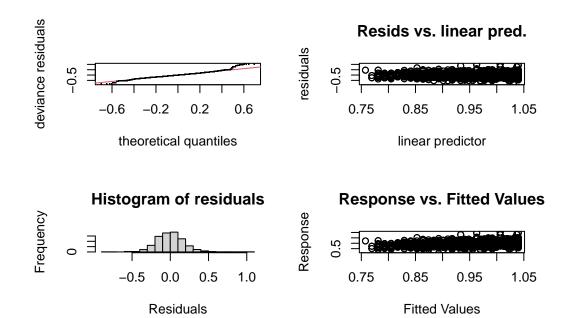
```
model <- gam(data = data_plot, WLK_TIME ~ s(AGE_VALS))
summary(model)</pre>
```

```
Family: gaussian
Link function: identity
Formula:
WLK_TIME ~ s(AGE_VALS)
Parametric coefficients:
           Estimate Std. Error t value Pr(>|t|)
(Intercept) 0.978068 0.001996 490.1 <2e-16 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Approximate significance of smooth terms:
             edf Ref.df
                           F p-value
s(AGE_VALS) 4.773 5.843 172.1 <2e-16 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
R-sq.(adj) = 0.109 Deviance explained = 10.9%
GCV = 0.032821 Scale est. = 0.032798 n = 8236
  plot(model)
```



Despite we don't have height values, it is possible to capture a decreasing trend of the walking speed with the age.

gam.check(model)



Method: GCV Optimizer: magic Smoothing parameter selection converged after 4 iterations. The RMS GCV score gradient at convergence was 4.021007e-07. The Hessian was positive definite. Model rank = 10 / 10

Basis dimension (k) checking results. Low p-value (k-index<1) may indicate that k is too low, especially if edf is close to k'.

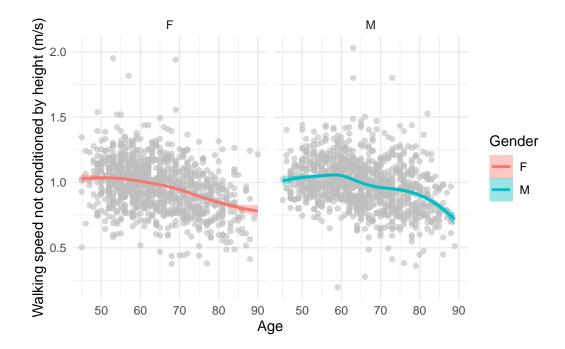
$$$\rm k'$$$
 edf k-index p-value s(AGE\_VALS) 9.00 4.77 1.01 0.67

To be continued...

If we split the data by gender:

```
geom_smooth(aes(color = SEX_ASK_COM, fill = SEX_ASK_COM)) +
facet_wrap(vars(SEX_ASK_COM)) +
xlab("Age") +
ylab("Walking speed not conditioned by height (m/s)") +
labs(fill = "Gender", color = "Gender") +
theme_minimal()
```

```
Joining with `by = join_by(id)`
`geom_smooth()` using method = 'gam' and formula = 'y ~ s(x, bs = "cs")'
```

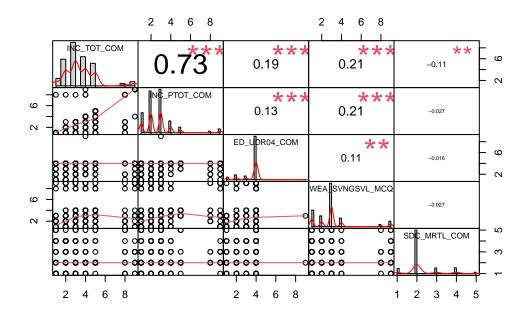


To analyze this data, I would add age and height as fixed effects using a generalized additive mixed model (GAMM). In (string2018?), the study is used as a random effect, but there were assessments across multiple study cases in different countries.

#### 3.3 Removed

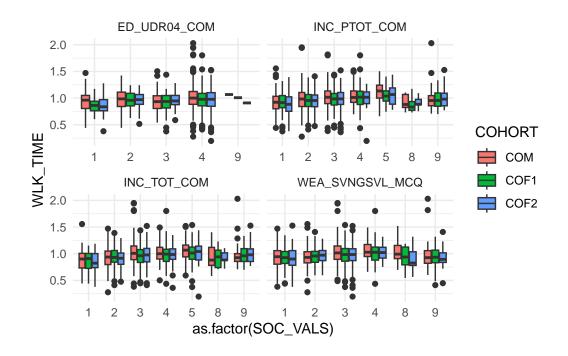
```
socvars <- data |>
  select(starts_with("INC"), ED_UDR04_COM, WEA_SVNGSVL_MCQ, SDC_MRTL_COM)
chart.Correlation(socvars, histogram=TRUE, pch=19)
```

```
Warning in par(usr): argument 1 does not name a graphical parameter
Warning in par(usr): argument 1 does not name a graphical parameter
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Warning in par(usr): argument 1 does not name a graphical parameter
Warning in par(usr): argument 1 does not name a graphical parameter
```



```
data_plot |>
ggplot( aes(x = as.factor(SOC_VALS), y = WLK_TIME)) +
  geom_boxplot(aes(fill = fct_relevel(COHORT, "COM", "COF1", "COF2"))) +
  facet_wrap(vars(SOC_VARS), scales = "free_x" ) +
  labs(fill = "COHORT") +
  theme_minimal()
```

Warning: Removed 1628 rows containing non-finite values (`stat\_boxplot()`).



```
# library(gamm4)
# model1 <- gamm(WLK_TIME ~ s(AGE_VALS) + INC_TOT_COM,</pre>
#
                  data = data_plot)
#
  model2 <- gamm(WLK_TIME ~ s(AGE_VALS) + INC_TOT_COM + ED_UDR04_COM + WEA_SVNGSVL_MCQ,</pre>
#
                  data = data_plot)
#
 model3 <- gamm(WLK_TIME ~ s(AGE_VALS, k = 4) +</pre>
                   s(INC_TOT_COM, k=4) +
#
                   s(ED\_UDRO4\_COM, k=4) +
                   s(WEA_SVNGSVL_MCQ, k=4),
#
                   data = data_soc)
#
#
#
# summary(model1)
# summary(model2)
# summary(model3)
```

## References

- Everitt, BS. 1998. "Analysis of Longitudinal Data: Beyond MANOVA." The British Journal of Psychiatry 172 (1): 7–10.
- Stringhini, Silvia, Cristian Carmeli, Markus Jokela, Mauricio Avendaño, Cathal McCrory, Angelo d'Errico, Murielle Bochud, et al. 2018. "Socioeconomic Status, Non-Communicable Disease Risk Factors, and Walking Speed in Older Adults: Multi-Cohort Population Based Study." *Bmj* 360.
- Twisk, Jos W. R. 2013. Applied Longitudinal Data Analysis for Epidemiology: A Practical Guide. 2nd ed. Cambridge University Press.