NEJM

```
library(tidyverse)
## -- Attaching packages ----- tidyverse 1.3.0 --
## v ggplot2 3.3.2
                    v purrr
                                0.3.4
## v tibble 3.0.4 v dplyr 1.0.2
## v tidyr 1.1.2 v stringr 1.4.0
## v readr 1.4.0 v forcats 0.5.0
## -- Conflicts ----- tidyverse_conflicts() --
## x dplyr::filter() masks stats::filter()
## x dplyr::lag() masks stats::lag()
library(dplyr)
library(tidyr)
library(knitr)
library(memisc)
## Loading required package: lattice
## Loading required package: MASS
## Attaching package: 'MASS'
## The following object is masked from 'package:dplyr':
##
##
       select
##
## Attaching package: 'memisc'
## The following objects are masked from 'package:dplyr':
##
##
       collect, recode, rename, syms
## The following object is masked from 'package:purrr':
##
##
       %@%
## The following object is masked from 'package:tibble':
##
##
       view
```

```
## The following object is masked from 'package:ggplot2':
##
##
       syms
## The following objects are masked from 'package:stats':
##
##
       contr.sum, contr.treatment, contrasts
## The following object is masked from 'package:base':
##
##
       as.array
library(qwraps2)
library(survival)
dig <- read_csv("dig.csv")</pre>
##
## -- Column specification -----
## cols(
##
     .default = col_double()
## )
## i Use 'spec()' for the full column specifications.
options(qwraps2_markup = "markdown")
#code applied from: https://cran.r-project.org/web/packages/qwraps2/vignettes/summary-statistics.html
#edited the n_perc0 function
get_p <- function (x, digits = getOption("qwraps2_frmt_digits", 1), na_rm = TRUE,</pre>
  show_denom = "never", show_symbol = TRUE, markup = getOption("qwraps2_markup",
    "latex"))
  d <- sum(!is.na(x))</pre>
  n <- sum(x, na.rm = na_rm)</pre>
  p <- frmt(100 * n/d, digits)
  if (show_denom == "never") {
   rtn <- paste0("", p)
  }
  if (!show_symbol) {
   rtn <- gsub("%", "", rtn)
  if (markup == "latex") {
    rtn <- gsub("%", "\\\\", rtn)
  }
  return(rtn)
our_summary1 <-
```

```
list(" " =
                                                     = ~ qwraps2::mean_sd(AGE, digits = 1),
        list("Age (yr) - mean ± SD"
             "Ejection fraction - mean ± SD"
                                                     = ~ qwraps2::mean_sd(EJF_PER, digits = 1),
             "Median duration of CHF - mo" = ~ median(CHFDUR, na.rm = T, digits = 1)),
      "% of patients" =
        list("Female sex" = ~ get_p(SEX == 2),
             "Nonwhite race" = ~ get_p(RACE == 2),
             "Age>70 yr" = \sim \text{get p(AGE} > 70)),
      "Method of assessing ejection fraction:" =
        list( "Radionuclide ventriculography" = ~ get_p(EJFMETH == 1),
              "Two-dimensional echocardiography" = ~ get_p(EJFMETH == 3),
              "Contrast angiography"= ~ get_p(EJFMETH == 2)),
      " " = list("Cardiothoracic ratio>0.55" =~ get_p(CHESTX > 0.55)),
      "NYHA class" = list("I" =~ get_p(FUNCTCLS == 1),
                           "II" =~ get_p(FUNCTCLS == 2),
                           "III" =~ get_p(FUNCTCLS == 3),
                           "IV"=~ get_p(FUNCTCLS == 4)
                           ),
      "No. of signs or symptoms of CHF†" = list("0" =~ get_p(NSYM == 0),
                                                  "1" =~ get_p(NSYM == 1),
                                                  "2" = \operatorname{get_p}(\operatorname{NSYM} == 2),
                                                  "3" =~ get_p(NSYM == 3),
                                                  "4+" = \operatorname{get_p(NSYM} > 3)),
      "Medical history" = list("Previous myocardial infarction" =~ get_p(PREVMI == 1),
                                "Current angina" =~ get_p(ANGINA == 1),
                                "Diabetes" =~ get_p(DIABETES == 1),
                                "Hypertension" =~ get p(HYPERTEN == 1)),
      " " = list("Previous digoxin use" =~ get_p(DIGUSE == 1)),
      "Primary Cause of Ischemia" = list( "Ischemic" =~ get_p(CHFETIOL == 1),
                                           "Nonischemic" =~ get_p(CHFETIOL != 1),
                                            "Nonischemic: idiopathic" =~ get_p(CHFETIOL == 4),
                                            "Nonischemic: hypertensive" =~ get_p(CHFETIOL == 2),
                                            "Nonischemic: other" =~ get_p(CHFETIOL %in% c(3,5,6))),
      "Concomitant medications" = list("Diuretics" =~ get_p(DIURET == 1 | DIURETK == 1),
                                         "Ace Inhibitors" =~ get_p(ACEINHIB == 1),
                                         "Nitrates" =~ get_p(NITRATES == 1),
                                        "Other Vasodilators" =~ get_p(VASOD == 1)),
      "Daily dose of study medication prescribed" = list("0.125 mg" =~ get_p(DIGDOSE == 0.125),
                                                           "0.250 \text{ mg"} = \text{get_p(DIGDOSE} == 0.25),
                                                           "0.375 \text{ mg}" = \text{get_p}(DIGDOSE} == 0.375),
                                                           "0.500 mg" =~ get_p(DIGDOSE == 0.5)))
summary_table(group_by(dig, TRTMT), our_summary1)
```

	0 (N = 3403)	1 (N = 3397)
** **		
Age (yr) — mean \pm SD	63.5 ± 10.8	63.4 ± 11.0
Ejection fraction — mean \pm SD	28.4 ± 8.9	28.6 ± 8.8
Median duration of CHF — mo	16	17
% of patients		
Female sex	22.5	22.2
Nonwhite race	14.8	14.3
Age > 70 yr	27.4	26.7

	0 (N = 3403)	1 (N = 3397)
Method of assessing ejection fraction:	/	
Radionuclide ventriculography	64.2	65.0
Two-dimensional echocardiography	30.0	29.5
Contrast angiography	5.8	5.5
**	0.0	0.0
Cardiothoracic ratio>0.55	34.4	34.6
NYHA class		
I	13.0	13.7
II	54.5	53.3
III	30.5	30.7
IV	1.9	2.2
No. of signs or symptoms of CHF†		
0	1.1	1.1
1	2.0	2.4
2	7.1	7.1
3	8.6	9.3
4+	81.2	80.2
Medical history		
Previous myocardial infarction	65.3	64.7
Current angina	26.4	27.1
Diabetes	28.6	28.3
Hypertension	45.8	45.0
**		
Previous digoxin use	44.6	44.1
Primary Cause of Ischemia		
Ischemic	70.7	71.0
Nonischemic	29.3	29.0
Nonischemic: idiopathic	14.2	15.5
Nonischemic: hypertensive	9.2	8.0
Nonischemic: other	6.0	5.5
Concomitant medications		
Diuretics	82.2	81.2
Ace Inhibitors	94.8	94.1
Nitrates	43.1	42.2
Other Vasodilators	1.5	0.9
Daily dose of study medication prescribe	\mathbf{d}	
0.125 mg	17.4	17.5
$0.250 \mathrm{\ mg}$	70.1	70.6
0.375 mg	11.3	10.3
$0.500 \mathrm{\ mg}$	0.9	1.1

Table 2

colnames(dig)

^{[1] &}quot;ID" "TRTMT" "AGE" "RACE" "SEX" "EJF_PER" [7] "EJFMETH" "CHESTX" "BMI" "KLEVEL" "CREAT" "DIGDOSER" [13] "CHFDUR" "RALES" "ELEVJVP" "PEDEMA" "RESTDYS" "EXERTDYS" [19] "ACTLIMIT" "S3" "PULCONG" "NSYM" "HEARTRTE" "DIABP"

^{[25] &}quot;SYSBP" "FUNCTCLS" "CHFETIOL" "PREVMI" "ANGINA" "DIABETES" [31] "HYPERTEN"

"DIGUSE" "DIURETK" "DIURET" "KSUPP" "ACEINHIB" [37] "NITRATES" "HYDRAL" "VASOD" "DIGDOSE" "CVD" "CVDDAYS" [43] "WHF" "WHFDAYS" "DIG" "DIGDAYS" "MI" "MIDAYS" [49] "UANG" "UANGDAYS" "STRK" "STRKDAYS" "SVA" "SVADAYS" [55] "VENA" "VENADAYS" "CREV" "CREVDAYS" "OCVD" "OCVDDAYS" [61] "RINF" "RINFDAYS" "OTH" "OTHDAYS" "HOSP" "HOSPDAYS" [67] "NHOSP" "DEATH" "DEATHDAY" "REASON" "DWHF" "DWHFDAYS"

	0 (N = 3403)	1 (N = 3397)
** **		
ALL	1,194 (35.1)	1,181 (34.8)
Cardiovascular	1,004 (29.5)	1,016 (29.9)
**		
Cardiovascular: Worsening Heart Failure	449 (37.6)	394 (33.4)
Cardiovascular: Other Cardiac	444 (37.2)	508 (43.0)
Cardiovascular: Other Vascular	45(3.8)	50 (4.2)
Cardiovascular: Unknown	66 (5.5)	64 (5.4)
Noncardiac and non-vascular	190 (15.9)	165 (14.0)

#COX Results

```
#cox models
dig <- dig %>%
  mutate(
    "death1" = case_when(
    REASON == 1 \sim 1,
    TRUE \sim 0),
    "death2" = case when(
    REASON == 2 \sim 1,
    TRUE \sim 0),
    "death3" = case_when(
    REASON == 3 \sim 1,
    TRUE \sim 0),
    "death4" = case_when(
    REASON == 4 \sim 1,
    TRUE \sim 0),
    "death5" = case_when(
    REASON == 5 \sim 1,
    TRUE \sim 0),
    "cardiodeath" = case_when(
      REASON \frac{1}{2} c(1:4) ~ 1,
```

```
TRUE ~ 0 )
    )
surva = Surv(dig$DEATHDAY, dig$DEATH)
survc = Surv(dig$DEATHDAY, dig$cardiodeath)
surv1=Surv(dig$DEATHDAY, dig$death1)
surv2=Surv(dig$DEATHDAY, dig$death2)
surv3=Surv(dig$DEATHDAY, dig$death3)
surv4=Surv(dig$DEATHDAY, dig$death4)
surv5=Surv(dig$DEATHDAY, dig$death5)
extract_p <- function(mod1){</pre>
 HR = exp(mod1$coefficients)
 lb = exp(confint(mod1))[1]
 ub = exp(confint(mod1))[2]
 p = summary(mod1)$coefficients[, 5]
 data.frame(HR, lb, ub, p)
}
mod1 <- coxph(surv1 ~ TRTMT, data = dig)</pre>
map_df(list(coxph(surva ~ TRTMT, data = dig),
            coxph(survc ~ TRTMT, data = dig),
            coxph(surv1 ~ TRTMT, data = dig),
            coxph(surv2 ~ TRTMT, data = dig),
            coxph(surv3 ~ TRTMT, data = dig),
            coxph(surv4 ~ TRTMT, data = dig),
            coxph(surv5 ~ TRTMT, data = dig)), ~extract_p(.x)) %>%
  knitr::kable()
```

	HR	lb	ub	p
$\overline{\text{TRTMT}1}$	0.9897291	0.9132349	1.072631	0.8013846
$\mathrm{TRTMT}\ldots 2$	1.0124118	0.9278499	1.104680	0.7816322
TRTMT3	0.8786338	0.7674446	1.005932	0.0608914
TRTMT4	1.1435411	1.0068172	1.298832	0.0389666
TRTMT5	1.1127996	0.7438911	1.664656	0.6029694
TRTMT6	0.9710935	0.6885412	1.369595	0.8672144
TRTMT7	0.8698142	0.7060665	1.071537	0.1899682