Data analysis

Signal detection of spontaneous medical device reports over time

Ty Stanford et al.

Table of contents

1	Set	up	2
	1.1	Packages	2
	1.2	Parallel compution setup	3
	1.3	Constants	
	1.4	Functions	4
	1.5	Load data	7
2	Ana	lysis	8
	2.1	BCPNN	8
	2.2	BCPNN with mult comp adjust	14
	2.3	PRR with mult comp adjust	21
	2.4	MaxSPRT	28
3	Rea	dy plot data	42
4	Sess	ion information	44

1 Set up

1.1 Packages

```
suppressPackageStartupMessages({
    library("readr")
    library("dplyr")
    library("tidyr")
    library("forcats")
    library("purrr")
    library("furrr")
    library("lubridate") # way to handle dates better than default R way
    library("tictoc")
                        # measure time elapsed in calcs
    library("ggplot2")
    library("ggrepel")
    library("knitr")
    library("gsDesign")
    library("foreach")
    library("arrow") # read/write parquet files
  })
Warning: package 'furrr' was built under R version 4.2.3
Warning: package 'future' was built under R version 4.2.3
Warning: package 'tictoc' was built under R version 4.2.3
Warning: package 'gsDesign' was built under R version 4.2.3
Warning: package 'foreach' was built under R version 4.2.3
  # NOTE: need to run first (only once, assumes devtools installed):
  # devtools::install_github("tystan/pharmsignal")
  library("pharmsignal") # signal detection algs
  # here are the functions written for these analyses
  # they will be shown in the *Appendix A*
  source("r/_funcs.R")
```

```
### NB: packages required that are used in above sourced file
# Sequential
# EmpiricalCalibration
```

1.2 Parallel compution setup

```
# options(future.globals.maxSize = 500 * 1024 ^ 2) # = 500 MiB
options(future.globals.maxSize = 2e3 * 1024 ^ 2) # = 2 GB

# furrr parallel workers/cores setup
# change `workers = 4` based on cores available in processor being used
(thread_to_use <- parallel::detectCores() - 2) # keep a core = 2 threads free</pre>
```

[1] 6

```
plan(multisession, workers = thread_to_use)

### test parallel works
# test code from https://furrr.futureverse.org/
# sequential
tic()
dev_null <- map(c(2, 2, 2), ~Sys.sleep(.x))
toc() # ~6 sec</pre>
```

6.14 sec elapsed

```
# parallel: should be (roughly, plus overheads) a third of the time of sequential
tic()
dev_null <- future_map(c(2, 2, 2), ~Sys.sleep(.x))
toc() # ~2 sec + overhead</pre>
```

4.14 sec elapsed

```
# for fun
tic()
dev_null <- future_map(rep(2, thread_to_use), ~Sys.sleep(.x))
toc()</pre>
```

4.28 sec elapsed

```
# this only applies to the non-parallel (non-"future") operations
set.seed(1234)
# this seed can be set in future_map() etc for reproducible parallel comp seeds
furrr_seed1 <- furrr_options(seed = 5678)
furrr_seed2 <- furrr_options(seed = 9012)
furrr_seed3 <- furrr_options(seed = 3456)</pre>
```

1.3 Constants

```
# arbitrarily, let's go with minimum cell count of 1 (will change based on context/applical arbitrary_cell_min <- 1
```

1.4 Functions

```
# do 90% CI only with lower == one sided 0.05
get_sig_tab <- function(nA, nB, nC, nD, alpha = 0.10, method = "bcpnn", n_mcmc = 1e+05) {
    out_cols_of_interest <- c("est_name", "est_scale", "est", "alpha", "ci_lo") # "ci_hi" (cols_tab <- NULL # initialise in scope
    if (method == "bcpnn") {
        sig_tab <- pharmsignal::bcpnn_mcmc_signal(nA, nB, nC, nD, alpha = alpha, n_mcmc = n_mc)
    } else if (method == "prr") {
        sig_tab <- pharmsignal::prr_signal(nA, nB, nC, nD, alpha = alpha)
    } else {
        stop("method for calcaultions unknown")
    }
    sig_tab <- sig_tab[, out_cols_of_interest]
    # sig_tab <- bind_cols(tibble(mnth = mnth), sig_tab)
    return(sig_tab)
}</pre>
```

```
get_sig_tab_over_time <- function(dat, alpha = 0.10, method = "bcpnn", n_mcmc = 1e+05) {</pre>
  n_tp <- nrow(dat)</pre>
  sig_tab_over_time <-</pre>
    foreach(i = 1:n_tp, .combine = bind_rows, .packages = "dplyr") %do% {
      with(
        dat,
        get_sig_tab(
          # mnth[i],
          nA[i], nB[i], nC[i], nD[i],
          alpha = alpha, method = method, n_mcmc = n_mcmc
        )
      )
    }
  return(sig_tab_over_time)
}
# if it's multiple comparisons central need to sparing use alpha
get_mult_compare_adj_alpha <- function(dat, alpha = 0.1) {</pre>
  n_reports <- nrow(dat)</pre>
  information_fracs <- (1:n_reports) / n_reports</pre>
  ### alternatives:
  # spend_obj <- sfLDPocock(alpha = alpha, t = information_fracs, param = NULL)
  \# spend_obj \leftarrow sfLDOF(alpha = alpha, t = information_fracs, param = NULL)
  spend_obj <- sfExponential(alpha = alpha, t = information_fracs, param = 0.5)</pre>
  # plot(1:n reports, spend_obj$spend, main = "alpha spending func", xlab = "look")
  return(bind_cols(dat, adj_alpha = spend_obj$spend))
}
```

```
# same as get_sig_tab_over_time(), however, alpha assumed included as column in data
  get_sig_tab_over_time_2 <- function(dat, method = "bcpnn", n_mcmc = 1e+05) {</pre>
    n_tp <- nrow(dat)</pre>
    sig_tab_over_time <-
      foreach(i = 1:n_tp, .combine = bind_rows, .packages = "dplyr") %do% {
        with(
          dat,
          get_sig_tab(
            # mnth[i],
            nA[i], nB[i], nC[i], nD[i],
            alpha = adj_alpha[i],
            method = method,
            n_mcmc = n_mcmc
          )
        )
      }
    return(sig_tab_over_time)
  }
  # test
  data.frame(nA = 30, nB = 5512, nC = 41, nD = 17445, adj_alpha = 0.1) %>%
    get_sig_tab_over_time_2(.)
   est_name est_scale
                             est alpha
                                           ci_lo
                  log2 0.7942907
                                   0.1 0.4317622
1 bcpnn mcmc
  data.frame(nA = 30, nB = 5512, nC = 41, nD = 17445, adj_alpha = 0.1) %>%
    get_sig_tab_over_time_2(., method = "prr")
 est_name est_scale
                           est alpha
                                        ci_lo
     prr orig scale 2.308667 0.1 1.556277
  2 ^ c(0.432304, 0.7942907) # similar to prr on ratio scale
[1] 1.349387 1.734225
```

```
log2(c(1.556277, 2.308667)) # similar to bcpnn on log2 scale
```

[1] 0.6380989 1.2070601

1.5 Load data

```
### monthly for testing
sra_dat <- read_parquet("dat/sra_dat.parquet")

### want this
cumul_qtrly_dat <- read_parquet("dat/cumul_qtrly_dat.parquet")

(thresholds <- sort(unique(sra_dat$thresh)))

[1] "0.010" "0.015" "0.020" "0.025" "0.030" "0.035" "0.040" "0.045" "0.050"
[10] "0.055" "0.060" "0.065" "0.070" "0.075" "0.080" "0.085" "0.090" "0.095"
[19] "0.100"</pre>
```

2 Analysis

2.1 BCPNN

```
sra_cum <-
    cumul_qtrly_dat
  # make data for each combination of params nested for purrr like processing
  sra_cum <-</pre>
    sra_cum %>%
    nest(data = c(mnth, nA, nB, nC, nD))
  sra_cum2 <-</pre>
    sra_dat %>%
    dplyr::filter(dat_type == "cumulative") %>%
    nest(data = c(mnth, nA, nB, nC, nD))
  # testing/example
  sra_cum$data[[9]] %>% print(., n = nrow(.))
# A tibble: 18 x 5
  mnth
              nA
                     nΒ
                           nC
                                  nD
   <chr>
           <dbl> <dbl> <dbl> <dbl> <dbl>
1 2013-Q3
               4
                     12
                                  10
2 2013-Q4
               6
                     14
                            1
                                  10
3 2014-Q1
               6
                     14
                                  11
                            1
4 2014-Q2
               7
                     15
                            1
                                  14
5 2014-Q3
               9
                     17
                            3
                                  21
6 2014-Q4
              26
                     19
                            4
                                  27
7 2015-Q1
              27
                     19
                                  28
              27
                                  28
8 2015-Q2
                     19
9 2015-Q3
              27
                     20
                                  28
10 2015-Q4
              27
                     20
                                  28
11 2016-Q1
              30
                                  28
                     21
                            6
12 2016-Q2
              34
                     21
                            6
                                  28
13 2016-Q3
              34
                     21
                            7
                                  33
14 2016-Q4
              36
                     23
                                  33
15 2017-Q1
              45
                     23
                            8
                                  34
                     24
                                  37
16 2017-Q2
              58
17 2017-Q3
              68
                     24
                                  38
```

18 2017-Q4 77 25 8 38

sra_cum2\$data[[9]] %>% print(., n = nrow(.))

#	A	tibble:	38 x	5		
		mnth	nA	nB	nC	nD
		<chr></chr>	<dbl></dbl>	<dbl></dbl>	<dbl></dbl>	<dbl></dbl>
1		2013-09	4	12	1	10
2	2	2013-11	6	13	1	10
3	3	2013-12	6	14	1	10
4		2014-03	6	14	1	11
5	•	2014-04	6	14	1	12
6	;	2014-05	7	15	1	13
7	•	2014-06	7	15	1	14
8	3	2014-07	7	15	2	15
9)	2014-08	9	17	2	19
10		2014-09	9	17	3	21
11		2014-10	10	18	3	24
12		2014-11	10	18	3	26
13		2014-12	26	19	4	27
14		2015-01	26	19	4	28
15		2015-03	27	19	4	28
16		2015-09	27	20	4	28
17		2015-10	27	20	5	28
18		2015-11	27	20	6	28
19		2016-01	29	20	6	28
20		2016-03	30	21	6	28
21		2016-04	34	21	6	28
22		2016-07	34	21	6	30
23		2016-08	34	21	7	32
24		2016-09	34	21	7	33
25		2016-11	36	22	7	33
26		2016-12	36	23	7	33
27		2017-01	40	23	7	33
28		2017-02	43	23	7	33
29		2017-03	45	23	8	34
30		2017-04	50	24	8	36
31		2017-05	54	24	8	37
32		2017-06	58	24	8	37
33		2017-07	60	24	8	38
34	Ŀ	2017-08	66	24	8	38

```
35 2017-09
              68
                    24
                           8
                                38
36 2017-10
              71
                                38
                    24
                           8
37 2017-11
              75
                    25
                           8
                                38
38 2017-12
              77
                    25
                           8
                                38
  get_sig_tab_over_time(sra_cum$data[[9]])
     est_name est_scale
                              est alpha
                                               ci_lo
                                    0.1 -0.27151962
                   log2 0.3778604
  bcpnn_mcmc
1
2 bcpnn_mcmc
                   log2 0.3738316
                                    0.1 -0.06911637
                                    0.1 -0.03108974
3 bcpnn_mcmc
                   log2 0.4150070
4 bcpnn_mcmc
                   log2 0.5127001
                                    0.1 0.11104715
                   log2 0.4951774
                                    0.1 0.06535270
5 bcpnn_mcmc
6 bcpnn_mcmc
                   log2 0.5370558
                                    0.1 0.33465530
7 bcpnn_mcmc
                   log2 0.5500767
                                    0.1 0.35025304
8 bcpnn_mcmc
                   log2 0.5500767
                                    0.1 0.35086730
                                    0.1 0.33836792
9 bcpnn_mcmc
                   log2 0.5377636
10 bcpnn_mcmc
                   log2 0.4850381
                                    0.1 0.28365055
11 bcpnn_mcmc
                   log2 0.4647627
                                    0.1 0.27878002
                   log2 0.4520794
                                    0.1 0.27929577
12 bcpnn_mcmc
13 bcpnn_mcmc
                   log2 0.5093946
                                    0.1 0.32733841
14 bcpnn_mcmc
                   log2 0.4823956
                                    0.1 0.30910135
15 bcpnn_mcmc
                   log2 0.4519206
                                    0.1 0.30245415
16 bcpnn_mcmc
                   log2 0.4402744
                                    0.1 0.30985519
17 bcpnn_mcmc
                   log2 0.4208998
                                    0.1 0.30189089
                   log2 0.3915001
                                    0.1 0.28170266
18 bcpnn mcmc
  ### for i5-8400/48GB 2133mhz memory
  # takes ~ 90 sec for monthly
  # takes ~ 40 sec for quarterly
  ### divide by a fair bit for r9-7900X
  tic()
  sra_cum <-
    sra_cum %>%
    mutate(
      sig_tab =
        future_map(
           .x = data,
           .f = get_sig_tab_over_time,
```

.options = furrr_seed1

```
)
  toc()
81.43 sec elapsed
  # check
  sra_cum$sig_tab[[9]]
     est_name est_scale
                              est alpha
                                               ci_lo
                                    0.1 -0.27857272
1 bcpnn_mcmc
                   log2 0.3778604
  bcpnn_mcmc
                   log2 0.3738316
                                    0.1 -0.06530536
2
                   log2 0.4150070
                                    0.1 -0.03003815
3 bcpnn_mcmc
4 bcpnn_mcmc
                   log2 0.5127001
                                    0.1 0.11613920
  bcpnn_mcmc
                   log2 0.4951774
                                    0.1
                                        0.06855644
                                    0.1 0.33396400
  bcpnn_mcmc
                   log2 0.5370558
7
  bcpnn_mcmc
                   log2 0.5500767
                                    0.1 0.34967589
                                    0.1 0.35032689
  bcpnn_mcmc
                   log2 0.5500767
  bcpnn_mcmc
                   log2 0.5377636
                                    0.1 0.33806521
10 bcpnn_mcmc
                   log2 0.4850381
                                    0.1 0.28283536
11 bcpnn_mcmc
                   log2 0.4647627
                                    0.1 0.27663489
12 bcpnn_mcmc
                   log2 0.4520794
                                    0.1 0.28009528
13 bcpnn_mcmc
                   log2 0.5093946
                                    0.1 0.33020338
14 bcpnn_mcmc
                   log2 0.4823956
                                    0.1 0.31108265
15 bcpnn_mcmc
                   log2 0.4519206
                                    0.1 0.30188642
16 bcpnn_mcmc
                   log2 0.4402744
                                    0.1 0.30898281
17 bcpnn_mcmc
                   log2 0.4208998
                                    0.1 0.30167104
18 bcpnn_mcmc
                   log2 0.3915001
                                    0.1 0.28215524
```

```
sra_cum_bcpnn <-
    sra_cum %>%
unnest(cols = c(data, sig_tab)) %>%
mutate(
    # dte = as_date(paste0(mnth, "-01"))
    dte =
        as_date(paste0(
            substr(mnth, 1, 5),
            sprintf("%02.0f", (as.integer(substr(mnth, 7, 7)) - 1) * 3 + 1),
            "-01"
        ))
```

```
)
  sra_cum_bcpnn
# A tibble: 1,707 x 14
  grps
              dat_t~1 thresh mnth
                                       nA
                                             nΒ
                                                   nC
                                                          nD est_n~2 est_s~3
                                                                                est
                                                                             <dbl>
   <chr>
                      <chr>
                             <chr> <dbl> <dbl> <dbl> <dbl> <chr>
                                                                     <chr>>
 1 (a) pelvi~ cumula~ 0.010
                             2013~
                                        3
                                              7
                                                    1
                                                           4 bcpnn_~ log2
                                                                             0.144
2 (a) pelvi~ cumula~ 0.010
                                                           5 bcpnn_~ log2
                             2013~
                                        4
                                             10
                                                    1
                                                                             0.170
3 (a) pelvi~ cumula~ 0.010
                             2013~
                                                    2
                                                           9 bcpnn_~ log2
                                        5
                                             11
                                                                             0.243
                                                           9 bcpnn_~ log2
4 (a) pelvi~ cumula~ 0.010
                             2013~
                                        9
                                             11
                                                    2
                                                                             0.323
5 (a) pelvi~ cumula~ 0.010 2014~
                                        9
                                             11
                                                    2
                                                          10 bcpnn_~ log2
                                                                             0.365
6 (a) pelvi~ cumula~ 0.010 2014~
                                       10
                                             12
                                                    3
                                                          12 bcpnn_~ log2
                                                                             0.351
7 (a) pelvi~ cumula~ 0.010 2014~
                                       12
                                                    5
                                                          19 bcpnn_~ log2
                                                                             0.420
                                             14
8 (a) pelvi~ cumula~ 0.010 2014~
                                                    7
                                                          24 bcpnn_~ log2
                                       30
                                             15
                                                                             0.445
9 (a) pelvi~ cumula~ 0.010 2015~
                                       31
                                             15
                                                    7
                                                          25 bcpnn_~ log2
                                                                             0.459
                                                    7
10 (a) pelvi~ cumula~ 0.010 2015~
                                       31
                                             15
                                                          25 bcpnn_~ log2
                                                                             0.459
# ... with 1,697 more rows, 3 more variables: alpha <dbl>, ci_lo <dbl>,
    dte <date>, and abbreviated variable names 1: dat_type, 2: est_name,
    3: est_scale
  # first signif
  bcpnn_signif <-
    sra_cum_bcpnn %>%
    group_by(grps, dat_type, thresh) %>%
    dplyr::filter(ci_lo > 0) %>%
    arrange(dte) %>%
    dplyr::filter(row_number() == 1) %>%
    ungroup() %>%
    rename(dte_reach_sig = dte)
  nrow(sra_cum_bcpnn)
[1] 1707
  sra_cum_bcpnn <-</pre>
    left_join(
      sra_cum_bcpnn,
```

```
bcpnn_signif %>% select(grps, dat_type, thresh, dte_reach_sig),
      c("grps", "dat_type", "thresh")
  nrow(sra_cum_bcpnn)
[1] 1707
  sra_cum_bcpnn
# A tibble: 1,707 x 15
              dat_t~1 thresh mnth
                                            nB
                                                  nC
                                                         nD est_n~2 est_s~3
  grps
                                      nA
                                                                              est
   <chr>
              <chr>
                      <chr>
                             <chr> <dbl> <dbl> <dbl> <dbl> <chr>
                                                                    <chr>
                                                                            <dbl>
                                                          4 bcpnn_~ log2
 1 (a) pelvi~ cumula~ 0.010
                             2013~
                                       3
                                             7
                                                    1
                                                                            0.144
2 (a) pelvi~ cumula~ 0.010 2013~
                                                          5 bcpnn_~ log2
                                                                            0.170
                                       4
                                             10
3 (a) pelvi~ cumula~ 0.010 2013~
                                                          9 bcpnn_~ log2
                                       5
                                            11
                                                    2
                                                                            0.243
4 (a) pelvi~ cumula~ 0.010 2013~
                                                    2
                                                          9 bcpnn_~ log2
                                       9
                                            11
                                                                            0.323
5 (a) pelvi~ cumula~ 0.010 2014~
                                       9
                                            11
                                                   2
                                                         10 bcpnn_~ log2
                                                                            0.365
6 (a) pelvi~ cumula~ 0.010 2014~
                                            12
                                                         12 bcpnn_~ log2
                                                                            0.351
                                      10
                                                   3
7 (a) pelvi~ cumula~ 0.010 2014~
                                                         19 bcpnn_~ log2
                                      12
                                            14
                                                    5
                                                                            0.420
                                                         24 bcpnn_~ log2
8 (a) pelvi~ cumula~ 0.010 2014~
                                      30
                                                    7
                                            15
                                                                            0.445
9 (a) pelvi~ cumula~ 0.010 2015~
                                      31
                                             15
                                                   7
                                                         25 bcpnn ~ log2
                                                                            0.459
10 (a) pelvi~ cumula~ 0.010 2015~
                                      31
                                             15
                                                    7
                                                         25 bcpnn_~ log2
                                                                            0.459
# ... with 1,697 more rows, 4 more variables: alpha <dbl>, ci_lo <dbl>,
    dte <date>, dte_reach_sig <date>, and abbreviated variable names
    1: dat_type, 2: est_name, 3: est_scale
  sra_cum_bcpnn <-</pre>
    sra_cum_bcpnn %>%
    mutate(
      dte_reach_sig = if_else(is.na(dte_reach_sig), as_date(today()), dte_reach_sig),
      reach_sig = dte >= dte_reach_sig
    )
  sra_cum_bcpnn %>%
    write_parquet(., sink = "out/sra_cum_bcpnn.parquet")
```

2.2 BCPNN with mult comp adjust

```
# sra_cum <-
  # sra_dat %>%
  # dplyr::filter(dat_type == "cumulative")
  sra_cum <-</pre>
    cumul_qtrly_dat
  sra_cum <-
    sra_cum %>%
    nest(data = c(mnth, nA, nB, nC, nD))
  # test get_mult_compare_adj_alpha()
  get_mult_compare_adj_alpha(sra_cum$data[[11]])
# A tibble: 18 x 6
  mnth
                   nВ
                         nC
                               nD adj_alpha
  <chr>
           <dbl> <dbl> <dbl> <dbl> <
                                      <dbl>
1 2013-Q3
              4
                   12
                               10 0.0000572
2 2013-Q4
                               10 0.001
              5
                   15
                          1
3 2014-Q1
              5
                   15
                             11 0.00355
                          1
4 2014-Q2
              6
                   16
                          1
                               14 0.00756
5 2014-Q3
             8
                   18
                          3 21 0.0127
6 2014-Q4
             25
                   20
                          4
                               27 0.0185
7 2015-Q1
             26
                   20
                               28 0.0249
                          4
8 2015-Q2
             26
                   20
                          4
                               28 0.0316
9 2015-Q3
             26
                   21
                          4 28 0.0385
10 2015-Q4
             26
                   21
                             30 0.0455
11 2016-Q1
             29
                   22
                             30 0.0526
                          4
12 2016-Q2
             33
                   22
                          4 30 0.0596
13 2016-Q3
                   22
                          5 35 0.0666
             33
14 2016-Q4
             35
                   24
                          5
                            35 0.0735
15 2017-Q1
             44
                   24
                          6 36 0.0803
                          6 39 0.0870
16 2017-Q2
             57
                   25
17 2017-Q3
                   25
                          6 40 0.0935
             67
                          6 40 0.1
18 2017-Q4
             76
                   26
```

get_sig_tab_over_time_2(get_mult_compare_adj_alpha(sra_cum\$data[[11]]))

```
est_name est_scale
                                          alpha
                                                     ci_lo
                              est
                   log2 0.3778604 5.719516e-05 -2.7775539
1
  bcpnn_mcmc
  bcpnn_mcmc
                   log2 0.3314299 1.000000e-03 -1.3130928
2
3
  bcpnn_mcmc
                   log2 0.3719247 3.552305e-03 -0.9735553
                   log2 0.4794164 7.562748e-03 -0.4878578
  bcpnn_mcmc
                   log2 0.4505463 1.266582e-02 -0.3737733
  bcpnn_mcmc
  bcpnn mcmc
                   log2 0.5291089 1.853315e-02
                                                 0.2334080
  bcpnn_mcmc
                   log2 0.5426635 2.491337e-02
                                                 0.2676446
  bcpnn_mcmc
                   log2 0.5426635 3.162278e-02
                                                 0.2787698
8
9
  bcpnn_mcmc
                   log2 0.5303613 3.852888e-02
                                                 0.2786349
10 bcpnn_mcmc
                   log2 0.5654356 4.553645e-02
                                                 0.3168146
11 bcpnn_mcmc
                   log2 0.5392350 5.257699e-02
                                                 0.3160579
                   log2 0.5198015 5.960122e-02
12 bcpnn_mcmc
                                                 0.3204447
13 bcpnn_mcmc
                   log2 0.5742360 6.657378e-02
                                                 0.3693870
14 bcpnn_mcmc
                   log2 0.5445855 7.346941e-02
                                                 0.3548644
15 bcpnn_mcmc
                   log2 0.5026231 8.027030e-02
                                                 0.3392577
16 bcpnn_mcmc
                   log2 0.4817174 8.696406e-02
                                                 0.3429426
17 bcpnn_mcmc
                   log2 0.4572070 9.354240e-02
                                                 0.3327737
                   log2 0.4241559 1.000000e-01
18 bcpnn_mcmc
                                                 0.3113718
```

get_sig_tab_over_time(sra_cum\$data[[11]])

```
est_name est_scale
                              est alpha
                                               ci_lo
                                    0.1 -0.27276457
                   log2 0.3778604
1 bcpnn mcmc
  bcpnn_mcmc
                   log2 0.3314299
                                    0.1 - 0.18935042
                   log2 0.3719247
                                    0.1 -0.15817042
  bcpnn mcmc
                                        0.01891085
  bcpnn_mcmc
                   log2 0.4794164
                   log2 0.4505463
                                    0.1 -0.02705216
5
  bcpnn_mcmc
6
  bcpnn_mcmc
                   log2 0.5291089
                                    0.1 0.32291402
7
  bcpnn_mcmc
                   log2 0.5426635
                                    0.1 0.33973052
8
  bcpnn_mcmc
                   log2 0.5426635
                                    0.1 0.33969117
                   log2 0.5303613
                                    0.1 0.32927472
  bcpnn_mcmc
10 bcpnn_mcmc
                   log2 0.5654356
                                    0.1 0.36001100
                                    0.1 0.34945147
11 bcpnn_mcmc
                   log2 0.5392350
12 bcpnn_mcmc
                   log2 0.5198015
                                    0.1
                                         0.34340898
                   log2 0.5742360
                                    0.1 0.38955969
13 bcpnn_mcmc
14 bcpnn mcmc
                   log2 0.5445855
                                    0.1 0.36988520
15 bcpnn_mcmc
                   log2 0.5026231
                                    0.1 0.34910679
```

```
16 bcpnn_mcmc
                    log2 0.4817174
                                      0.1 0.34742783
                    log2 0.4572070
                                      0.1 0.33516223
17 bcpnn_mcmc
18 bcpnn_mcmc
                    log2 0.4241559
                                      0.1 0.31176932
  tic()
  sra_cum <-</pre>
    sra_cum %>%
    mutate(
      data =
        map(
          .x = data,
          .f = get_mult_compare_adj_alpha
        )
    )
  toc()
```

0.17 sec elapsed

```
# test
sra_cum$data[[11]] # check adj_alpha added as column in data
```

```
# A tibble: 18 x 6
  mnth
               nA
                     nВ
                            nC
                                  nD adj_alpha
   <chr>>
                                          <dbl>
           <dbl> <dbl> <dbl> <dbl> <
1 2013-Q3
                4
                     12
                             1
                                  10 0.0000572
2 2013-Q4
                5
                     15
                             1
                                  10 0.001
3 2014-Q1
                5
                     15
                                  11 0.00355
4 2014-Q2
                6
                     16
                                  14 0.00756
                             1
5 2014-Q3
                8
                     18
                             3
                                  21 0.0127
6 2014-Q4
               25
                     20
                             4
                                  27 0.0185
7 2015-Q1
               26
                     20
                                  28 0.0249
               26
8 2015-Q2
                     20
                                  28 0.0316
9 2015-Q3
               26
                     21
                             4
                                  28 0.0385
10 2015-Q4
               26
                     21
                                  30 0.0455
11 2016-Q1
               29
                     22
                             4
                                  30 0.0526
12 2016-Q2
               33
                     22
                             4
                                  30 0.0596
13 2016-Q3
                     22
               33
                             5
                                  35 0.0666
14 2016-Q4
               35
                     24
                             5
                                  35 0.0735
15 2017-Q1
               44
                     24
                             6
                                  36 0.0803
16 2017-Q2
               57
                     25
                                  39 0.0870
```

```
18 2017-Q4
                                 40 0.1
              76
                    26
                            6
  ### takes ~ 100 sec (i5-8400 6c/6t)
  ### takes ~55 sec on laptop lols (i5 8th gen 4c/8t)
  tic()
  sra_cum <-
    sra_cum %>%
    mutate(
      sig_tab =
        future_map(
           .x = data,
          .f = get_sig_tab_over_time_2, # the alpha in data version
           .options = furrr_seed1
        )
    )
  toc()
```

40 0.0935

83.12 sec elapsed

17 2017-Q3

67

25

6

```
# check
sra_cum$sig_tab[[11]]
```

```
est_name est_scale
                                         alpha
                              est
                                                    ci_lo
 bcpnn_mcmc
                   log2 0.3778604 5.719516e-05 -2.4252741
2 bcpnn_mcmc
                   log2 0.3314299 1.000000e-03 -1.2692260
3 bcpnn_mcmc
                   log2 0.3719247 3.552305e-03 -0.9503514
                   log2 0.4794164 7.562748e-03 -0.4816096
4 bcpnn_mcmc
5 bcpnn_mcmc
                   log2 0.4505463 1.266582e-02 -0.3786932
6 bcpnn_mcmc
                   log2 0.5291089 1.853315e-02 0.2341781
7 bcpnn_mcmc
                   log2 0.5426635 2.491337e-02 0.2692897
8 bcpnn mcmc
                   log2 0.5426635 3.162278e-02 0.2796775
9 bcpnn_mcmc
                   log2 0.5303613 3.852888e-02
                                               0.2788266
10 bcpnn mcmc
                   log2 0.5654356 4.553645e-02 0.3196894
11 bcpnn_mcmc
                   log2 0.5392350 5.257699e-02 0.3167301
12 bcpnn_mcmc
                   log2 0.5198015 5.960122e-02 0.3204761
13 bcpnn_mcmc
                   log2 0.5742360 6.657378e-02 0.3686319
                   log2 0.5445855 7.346941e-02 0.3541094
14 bcpnn_mcmc
                   log2 0.5026231 8.027030e-02
15 bcpnn_mcmc
                                               0.3392261
16 bcpnn_mcmc
                   log2 0.4817174 8.696406e-02 0.3425696
```

```
17 bcpnn_mcmc
                   log2 0.4572070 9.354240e-02 0.3331946
18 bcpnn_mcmc
                   log2 0.4241559 1.000000e-01 0.3126421
  sra_cum_bcpnn_mc_adj <-</pre>
    sra_cum %>%
    unnest(cols = c(data, sig_tab)) %>%
    mutate(
      # dte = as date(pasteO(mnth, "-01"))
      dte =
        as date(paste0(
          substr(mnth, 1, 5),
          sprintf(\%02.0f'', (as.integer(substr(mnth, 7, 7)) - 1) * 3 + 1),
          "-01"
        ))
    )
  sra_cum_bcpnn_mc_adj
# A tibble: 1,707 x 15
            dat_t~1 thresh mnth
                                                nC
                                                      nD adj_a~2 est_n~3 est_s~4
                                    nA
                                          nΒ
  grps
   <chr>
                           <chr> <dbl> <dbl> <dbl> <dbl>
                                                            <dbl> <chr>
            <chr>
                    <chr>
 1 (a) pel~ cumula~ 0.010 2013~
                                     3
                                           7
                                                  1
                                                        4 3.37e-5 bcpnn_~ log2
2 (a) pel~ cumula~ 0.010 2013~
                                          10
                                                        5 6.88e-4 bcpnn_~ log2
3 (a) pel~ cumula~ 0.010 2013~
                                     5
                                          11
                                                 2
                                                        9 2.62e-3 bcpnn_~ log2
                                                       9 5.81e-3 bcpnn_~ log2
4 (a) pel~ cumula~ 0.010 2013~
                                                 2
                                     9
                                          11
5 (a) pel~ cumula~ 0.010 2014~
                                     9
                                          11
                                                 2
                                                       10 1
                                                              e-2 bcpnn_~ log2
6 (a) pel~ cumula~ 0.010 2014~
                                          12
                                                 3
                                                       12 1.49e-2 bcpnn ~ log2
                                    10
7 (a) pel~ cumula~ 0.010 2014~
                                    12
                                          14
                                                 5
                                                       19 2.04e-2 bcpnn_~ log2
8 (a) pel~ cumula~ 0.010 2014~
                                    30
                                          15
                                                 7
                                                       24 2.62e-2 bcpnn ~ log2
9 (a) pel~ cumula~ 0.010 2015~
                                    31
                                          15
                                                 7
                                                       25 3.23e-2 bcpnn_~ log2
                                                 7
10 (a) pel~ cumula~ 0.010 2015~
                                    31
                                          15
                                                       25 3.85e-2 bcpnn_~ log2
# ... with 1,697 more rows, 4 more variables: est <dbl>, alpha <dbl>,
    ci lo <dbl>, dte <date>, and abbreviated variable names 1: dat type,
    2: adj_alpha, 3: est_name, 4: est_scale
  # first signif
  bcpnn_mc_adj_signif <-
    sra_cum_bcpnn_mc_adj %>%
    group_by(grps, dat_type, thresh) %>%
    dplyr::filter(ci_lo > 0) %>%
```

```
arrange(dte) %>%
    dplyr::filter(row_number() == 1) %>%
    ungroup() %>%
    rename(dte_reach_sig = dte)
  nrow(sra_cum_bcpnn_mc_adj)
[1] 1707
  sra_cum_bcpnn_mc_adj <-</pre>
    left_join(
      sra_cum_bcpnn_mc_adj,
      bcpnn_mc_adj_signif %>% select(grps, dat_type, thresh, dte_reach_sig),
      c("grps", "dat_type", "thresh")
    )
  nrow(sra_cum_bcpnn_mc_adj)
[1] 1707
  sra_cum_bcpnn_mc_adj
# A tibble: 1,707 x 16
  grps
            dat t~1 thresh mnth
                                                       nD adj a~2 est n~3 est s~4
                                    nA
                                          nВ
                                                 nC
            <chr>
                                                            <dbl> <chr>
                    <chr> <chr> <dbl> <dbl> <dbl> <dbl> <dbl>
 1 (a) pel~ cumula~ 0.010 2013~
                                     3
                                           7
                                                  1
                                                        4 3.37e-5 bcpnn_~ log2
2 (a) pel~ cumula~ 0.010 2013~
                                                        5 6.88e-4 bcpnn_~ log2
                                     4
                                           10
                                                  1
3 (a) pel~ cumula~ 0.010 2013~
                                     5
                                           11
                                                  2
                                                        9 2.62e-3 bcpnn_~ log2
4 (a) pel~ cumula~ 0.010 2013~
                                     9
                                           11
                                                  2
                                                        9 5.81e-3 bcpnn_~ log2
5 (a) pel~ cumula~ 0.010 2014~
                                     9
                                                  2
                                                              e-2 bcpnn_~ log2
                                           11
                                                       10 1
6 (a) pel~ cumula~ 0.010 2014~
                                    10
                                           12
                                                  3
                                                       12 1.49e-2 bcpnn_~ log2
7 (a) pel~ cumula~ 0.010 2014~
                                    12
                                                       19 2.04e-2 bcpnn_~ log2
                                           14
                                                  7
8 (a) pel~ cumula~ 0.010 2014~
                                    30
                                           15
                                                       24 2.62e-2 bcpnn_~ log2
9 (a) pel~ cumula~ 0.010 2015~
                                    31
                                           15
                                                  7
                                                       25 3.23e-2 bcpnn_~ log2
                                                       25 3.85e-2 bcpnn_~ log2
10 (a) pel~ cumula~ 0.010 2015~
                                    31
                                           15
                                                  7
# ... with 1,697 more rows, 5 more variables: est <dbl>, alpha <dbl>,
   ci_lo <dbl>, dte <date>, dte_reach_sig <date>, and abbreviated variable
   names 1: dat_type, 2: adj_alpha, 3: est_name, 4: est_scale
```

```
sra_cum_bcpnn_mc_adj <-
    sra_cum_bcpnn_mc_adj %>%
mutate(
    dte_reach_sig = if_else(is.na(dte_reach_sig), as_date(today()), dte_reach_sig),
    reach_sig = dte >= dte_reach_sig
)

sra_cum_bcpnn_mc_adj %>%
write_parquet(., sink = "out/sra_cum_bcpnn_mc_adj.parquet")
```

2.3 PRR with mult comp adjust

```
# sra_cum <-
  # sra_dat %>%
  # dplyr::filter(dat_type == "cumulative")
  sra_cum <-</pre>
    cumul_qtrly_dat
  sra_cum <-
    sra_cum %>%
    nest(data = c(mnth, nA, nB, nC, nD))
  # test
  get_mult_compare_adj_alpha(sra_cum$data[[11]])
# A tibble: 18 x 6
  mnth
                   nΒ
                         nC
                               nD adj_alpha
   <chr>
           <dbl> <dbl> <dbl> <dbl> <
                                      <dbl>
1 2013-Q3
              4
                               10 0.0000572
                   12
                               10 0.001
2 2013-Q4
              5
                   15
                          1
3 2014-Q1
              5
                   15
                             11 0.00355
                          1
4 2014-Q2
              6
                   16
                          1
                               14 0.00756
5 2014-Q3
             8
                   18
                          3 21 0.0127
6 2014-Q4
              25
                   20
                          4
                               27 0.0185
7 2015-Q1
              26
                   20
                               28 0.0249
                          4
8 2015-Q2
              26
                    20
                          4
                               28 0.0316
9 2015-Q3
              26
                   21
                          4
                               28 0.0385
10 2015-Q4
              26
                   21
                             30 0.0455
11 2016-Q1
              29
                    22
                          4
                               30 0.0526
12 2016-Q2
              33
                   22
                          4 30 0.0596
13 2016-Q3
                    22
                          5
              33
                               35 0.0666
14 2016-Q4
              35
                   24
                          5
                               35 0.0735
15 2017-Q1
              44
                   24
                          6 36 0.0803
                          6 39 0.0870
16 2017-Q2
              57
                    25
17 2017-Q3
                          6 40 0.0935
              67
                    25
                          6 40 0.1
18 2017-Q4
             76
                    26
```

```
get_sig_tab_over_time_2(get_mult_compare_adj_alpha(sra_cum$data[[11]]))
```

est

est_name est_scale

```
log2 0.3778604 5.719516e-05 -2.3823226
1
 bcpnn_mcmc
  bcpnn_mcmc
                   log2 0.3314299 1.000000e-03 -1.3454188
2
3
  bcpnn_mcmc
                   log2 0.3719247 3.552305e-03 -0.9331261
  bcpnn mcmc
                   log2 0.4794164 7.562748e-03 -0.4892387
  bcpnn_mcmc
                   log2 0.4505463 1.266582e-02 -0.3800232
6
  bcpnn mcmc
                   log2 0.5291089 1.853315e-02 0.2342780
  bcpnn_mcmc
                   log2 0.5426635 2.491337e-02
                                                0.2666768
 bcpnn_mcmc
                   log2 0.5426635 3.162278e-02
                                                0.2782165
8
  bcpnn_mcmc
                   log2 0.5303613 3.852888e-02
                                                0.2785088
10 bcpnn_mcmc
                   log2 0.5654356 4.553645e-02
                                                0.3178754
11 bcpnn_mcmc
                   log2 0.5392350 5.257699e-02
                                                0.3159453
                   log2 0.5198015 5.960122e-02
                                                0.3207910
12 bcpnn_mcmc
13 bcpnn_mcmc
                   log2 0.5742360 6.657378e-02
                                                0.3682926
14 bcpnn_mcmc
                   log2 0.5445855 7.346941e-02
                                                0.3535223
15 bcpnn_mcmc
                   log2 0.5026231 8.027030e-02
                                                0.3395410
16 bcpnn_mcmc
                   log2 0.4817174 8.696406e-02
                                                0.3430154
17 bcpnn_mcmc
                   log2 0.4572070 9.354240e-02
                                                0.3327655
                   log2 0.4241559 1.000000e-01
18 bcpnn_mcmc
                                                0.3124256
  get_sig_tab_over_time_2(get_mult_compare_adj_alpha(sra_cum$data[[11]]), method = "prr")
```

alpha

ci lo

```
est_name
             est_scale
                                        alpha
                                                   ci lo
                            est
1
        prr orig scale 2.750000 5.719516e-05 0.04066616
2
        prr orig scale 2.750000 1.000000e-03 0.09303769
3
        prr orig scale 3.000000 3.552305e-03 0.14772511
4
        prr orig scale 4.090909 7.562748e-03 0.26339441
5
        prr orig scale 2.461538 1.266582e-02 0.53132229
6
        prr orig scale 4.305556 1.853315e-02 1.37316395
7
        prr orig scale 4.521739 2.491337e-02 1.52286634
8
        prr orig scale 4.521739 3.162278e-02 1.59362698
9
        prr orig scale 4.425532 3.852888e-02 1.61983333
10
        prr orig scale 4.702128 4.553645e-02 1.77347101
11
        prr orig scale 4.833333 5.257699e-02 1.88692159
12
        prr orig scale 5.100000 5.960122e-02 2.05551981
13
        prr orig scale 4.800000 6.657378e-02 2.17067746
14
        prr orig scale 4.745763 7.346941e-02 2.19025371
15
        prr orig scale 4.529412 8.027030e-02 2.29602631
```

```
16
        prr orig scale 5.213415 8.696406e-02 2.68795062
17
        prr orig scale 5.583333 9.354240e-02 2.92265162
18
        prr orig scale 5.712418 1.000000e-01 3.03209290
  get_sig_tab_over_time(sra_cum$data[[11]], method = "prr")
                             est alpha
                                           ci_lo
   est_name est_scale
        prr orig scale 2.750000
                                   0.1 0.4912212
1
2
        prr orig scale 2.750000
                                   0.1 0.5060331
3
        prr orig scale 3.000000
                                   0.1 0.5487100
        prr orig scale 4.090909
4
                                   0.1 0.7555189
5
        prr orig scale 2.461538
                                   0.1 0.8951429
        prr orig scale 4.305556
6
                                   0.1 1.9379924
7
        prr orig scale 4.521739
                                   0.1 2.0354629
8
        prr orig scale 4.521739
                                   0.1 2.0354629
        prr orig scale 4.425532
                                   0.1 1.9905948
9
10
        prr orig scale 4.702128
                                   0.1 2.1084471
        prr orig scale 4.833333
                                   0.1 2.1757613
11
12
        prr orig scale 5.100000
                                   0.1 2.3065749
13
        prr orig scale 4.800000
                                   0.1 2.3563014
14
        prr orig scale 4.745763
                                   0.1 2.3318890
15
        prr orig scale 4.529412
                                   0.1 2.3909246
        prr orig scale 5.213415
                                   0.1 2.7583297
16
17
        prr orig scale 5.583333
                                   0.1 2.9591422
18
        prr orig scale 5.712418
                                   0.1 3.0320929
  tic()
  sra_cum <-</pre>
    sra_cum %>%
    mutate(
      data =
        map(
          .x = data,
           .f = get_mult_compare_adj_alpha
        )
    )
  toc()
```

0.17 sec elapsed

```
# test
  sra_cum$data[[11]] # check adj_alpha added as column in data
# A tibble: 18 x 6
              nA
  mnth
                    nΒ
                          nC
                                 nD adj_alpha
           <dbl> <dbl> <dbl> <dbl> <
                                        <dbl>
   <chr>>
1 2013-Q3
               4
                    12
                            1
                                 10 0.0000572
2 2013-Q4
               5
                    15
                            1
                                 10 0.001
3 2014-Q1
               5
                    15
                                 11 0.00355
                            1
4 2014-Q2
               6
                    16
                           1
                                14 0.00756
5 2014-Q3
               8
                    18
                           3
                                 21 0.0127
6 2014-Q4
              25
                    20
                           4
                                27 0.0185
7 2015-Q1
              26
                    20
                           4
                                 28 0.0249
8 2015-Q2
                    20
                                28 0.0316
              26
                           4
9 2015-Q3
              26
                                28 0.0385
                    21
                           4
10 2015-Q4
              26
                    21
                                 30 0.0455
              29
11 2016-Q1
                    22
                                30 0.0526
12 2016-Q2
              33
                    22
                           4
                                30 0.0596
13 2016-Q3
                              35 0.0666
              33
                    22
                           5
14 2016-Q4
              35
                    24
                           5
                                 35 0.0735
15 2017-Q1
              44
                    24
                           6 36 0.0803
16 2017-Q2
              57
                    25
                           6
                                 39 0.0870
                              40 0.0935
17 2017-Q3
              67
                    25
                           6
18 2017-Q4
              76
                    26
                                 40 0.1
  get_sig_tab_over_time_2_prr <- function(dat) {</pre>
   get_sig_tab_over_time_2(dat, method = "prr")
  }
  ### takes ~2 sec on laptop (i5 8th gen 4c/8t)
  tic()
  sra_cum <-</pre>
    sra_cum %>%
    mutate(
      sig_tab =
        future_map(
          .x = data,
          .f = get_sig_tab_over_time_2_prr, # the alpha in data version
           .options = furrr_seed1
        )
```

```
)
toc()
```

))

```
3.37 sec elapsed
  # check
  sra_cum$sig_tab[[11]]
   est_name est_scale
                             est
                                        alpha
                                                   ci lo
1
        prr orig scale 2.750000 5.719516e-05 0.04066616
2
        prr orig scale 2.750000 1.000000e-03 0.09303769
3
        prr orig scale 3.000000 3.552305e-03 0.14772511
4
        prr orig scale 4.090909 7.562748e-03 0.26339441
5
        prr orig scale 2.461538 1.266582e-02 0.53132229
6
        prr orig scale 4.305556 1.853315e-02 1.37316395
7
        prr orig scale 4.521739 2.491337e-02 1.52286634
8
        prr orig scale 4.521739 3.162278e-02 1.59362698
9
        prr orig scale 4.425532 3.852888e-02 1.61983333
10
        prr orig scale 4.702128 4.553645e-02 1.77347101
11
        prr orig scale 4.833333 5.257699e-02 1.88692159
12
        prr orig scale 5.100000 5.960122e-02 2.05551981
13
        prr orig scale 4.800000 6.657378e-02 2.17067746
14
        prr orig scale 4.745763 7.346941e-02 2.19025371
15
        prr orig scale 4.529412 8.027030e-02 2.29602631
16
        prr orig scale 5.213415 8.696406e-02 2.68795062
17
        prr orig scale 5.583333 9.354240e-02 2.92265162
18
        prr orig scale 5.712418 1.000000e-01 3.03209290
  sra_cum_prr_mc_adj <-</pre>
    sra_cum %>%
    unnest(cols = c(data, sig_tab)) %>%
    mutate(
      # dte = as_date(pasteO(mnth, "-01"))
      dte =
        as_date(paste0(
          substr(mnth, 1, 5),
          sprintf(\%02.0f'', (as.integer(substr(mnth, 7, 7)) - 1) * 3 + 1),
          "-01"
```

```
)
  sra_cum_prr_mc_adj
# A tibble: 1,707 x 15
  grps
            dat_t~1 thresh mnth
                                    nA
                                           nВ
                                                 nC
                                                       nD adj_a~2 est_n~3 est_s~4
                                                            <dbl> <chr>
   <chr>
                    <chr> <chr> <dbl> <dbl> <dbl> <dbl> <
                                                                           <chr>
 1 (a) pel~ cumula~ 0.010 2013~
                                     3
                                            7
                                                  1
                                                        4 3.37e-5 prr
                                                                           orig s~
2 (a) pel~ cumula~ 0.010 2013~
                                           10
                                                        5 6.88e-4 prr
                                      4
                                                  1
                                                                           orig s~
3 (a) pel~ cumula~ 0.010 2013~
                                     5
                                                  2
                                                        9 2.62e-3 prr
                                                                           orig s~
                                           11
4 (a) pel~ cumula~ 0.010 2013~
                                                  2
                                     9
                                           11
                                                        9 5.81e-3 prr
                                                                           orig s~
5 (a) pel~ cumula~ 0.010 2014~
                                                  2
                                     9
                                           11
                                                       10 1
                                                              e-2 prr
                                                                           orig s~
6 (a) pel~ cumula~ 0.010 2014~
                                    10
                                           12
                                                  3
                                                       12 1.49e-2 prr
                                                                           orig s~
7 (a) pel~ cumula~ 0.010 2014~
                                           14
                                                  5
                                                       19 2.04e-2 prr
                                    12
                                                                           orig s~
8 (a) pel~ cumula~ 0.010 2014~
                                                  7
                                    30
                                           15
                                                       24 2.62e-2 prr
                                                                           orig s~
9 (a) pel~ cumula~ 0.010 2015~
                                    31
                                           15
                                                  7
                                                       25 3.23e-2 prr
                                                                           orig s~
10 (a) pel~ cumula~ 0.010 2015~
                                    31
                                           15
                                                  7
                                                       25 3.85e-2 prr
                                                                           orig s~
# ... with 1,697 more rows, 4 more variables: est <dbl>, alpha <dbl>,
   ci_lo <dbl>, dte <date>, and abbreviated variable names 1: dat_type,
    2: adj_alpha, 3: est_name, 4: est_scale
  # first signif
  prr_mc_adj_signif <-</pre>
    sra_cum_prr_mc_adj %>%
    group_by(grps, dat_type, thresh) %>%
    dplyr::filter(ci_lo > 1) %>% # 1 is the critical value on ratio scale
    arrange(dte) %>%
    dplyr::filter(row_number() == 1) %>%
    ungroup() %>%
    rename(dte_reach_sig = dte)
  nrow(sra_cum_prr_mc_adj)
[1] 1707
  sra_cum_prr_mc_adj <-</pre>
    left_join(
      sra_cum_prr_mc_adj,
```

```
prr_mc_adj_signif %>% select(grps, dat_type, thresh, dte_reach_sig),
      c("grps", "dat_type", "thresh")
  nrow(sra_cum_prr_mc_adj)
[1] 1707
  sra_cum_prr_mc_adj
# A tibble: 1,707 x 16
           dat_t~1 thresh mnth
                                                nC
                                                      nD adj_a~2 est_n~3 est_s~4
  grps
                                   nA
                                          nВ
  <chr>
            <chr>
                   <chr> <chr> <dbl> <dbl> <dbl> <dbl> <dbl>
                                                           <dbl> <chr>
                                                                         <chr>
 1 (a) pel~ cumula~ 0.010 2013~
                                     3
                                           7
                                                       4 3.37e-5 prr
                                                 1
                                                                         orig s~
2 (a) pel~ cumula~ 0.010 2013~
                                          10
                                                 1
                                                       5 6.88e-4 prr
                                                                         orig s~
                                     4
3 (a) pel~ cumula~ 0.010 2013~
                                     5
                                          11
                                                 2
                                                       9 2.62e-3 prr
                                                                         orig s~
4 (a) pel~ cumula~ 0.010 2013~
                                                 2
                                     9
                                         11
                                                       9 5.81e-3 prr
                                                                         orig s~
5 (a) pel~ cumula~ 0.010 2014~
                                    9
                                          11
                                                 2
                                                      10 1
                                                             e-2 prr
                                                                         orig s~
6 (a) pel~ cumula~ 0.010 2014~
                                          12
                                                 3
                                                      12 1.49e-2 prr
                                                                         orig s~
                                    10
7 (a) pel~ cumula~ 0.010 2014~
                                    12
                                          14
                                                 5
                                                      19 2.04e-2 prr
                                                                         orig s~
8 (a) pel~ cumula~ 0.010 2014~
                                                 7
                                                      24 2.62e-2 prr
                                    30
                                          15
                                                                         orig s~
9 (a) pel~ cumula~ 0.010 2015~
                                                 7
                                                      25 3.23e-2 prr
                                    31
                                          15
                                                                         orig s~
10 (a) pel~ cumula~ 0.010 2015~
                                    31
                                          15
                                                 7
                                                      25 3.85e-2 prr
                                                                         orig s~
# ... with 1,697 more rows, 5 more variables: est <dbl>, alpha <dbl>,
   ci_lo <dbl>, dte <date>, dte_reach_sig <date>, and abbreviated variable
   names 1: dat_type, 2: adj_alpha, 3: est_name, 4: est_scale
  sra_cum_prr_mc_adj <-</pre>
    sra_cum_prr_mc_adj %>%
    mutate(
      dte_reach_sig = if_else(is.na(dte_reach_sig), as_date(today()), dte_reach_sig),
      reach_sig = dte >= dte_reach_sig
    )
  sra_cum_prr_mc_adj %>%
    write_parquet(., sink = "out/sra_cum_prr_mc_adj.parquet")
```

2.4 MaxSPRT

```
# sra_cum <-
# sra_dat %>%
# dplyr::filter(dat_type == "cumulative")
sra_cum <-</pre>
 cumul_qtrly_dat
cv_tab <-
  sra_cum %>%
  # dplyr::filter(thresh < 0.070) %>%
  group_by(grps, thresh) %>%
  summarise(
   min_dte = min(mnth),
   \max_{dte} = \max_{dte}(mnth),
   rows = n(),
   sum_nA = max(nA),
   sum_nC = max(nC),
   tot_n = sum_nA + sum_nC,
    .groups = "drop"
  ) %>%
  mutate(
    \# qtrs = interval(paste0(min_dte, "-01"), paste0(max_dte, "-01")) / months(1) / 4,
   qtrs = rows,
  n_per_qtr = tot_n / qtrs,
   z = sum_nC / sum_nA
  )
cv_tab %>%
 kable(., digits = 1)
```

grps	threshmi	n_d te ax_	_d rte ws	sum_	_nsAm_	_nt6t	nqtrs	n_per_	_qt z
(a) pelvic_mesh v	0.010 20	13- 2017-	- 20	82	12	94	20	4.7	0.1
hernia_mesh	Q_1	Q4							
(a) pelvic_mesh v	0.015 20	13- 2017-	- 20	82	12	94	20	4.7	0.1
hernia_mesh	Q_1	Q4							
(a) pelvic_mesh v	0.020 20	13- 2017-	- 18	82	10	92	18	5.1	0.1
hernia_mesh	Q:	Q4							
(a) pelvic_mesh v	0.025 20	13- 2017-	- 18	82	10	92	18	5.1	0.1
hernia mesh	Q:	Q4							

grps	thresh	nmin_	_dtmax_	_dtews	sum_	nsAm_	_nt6t_	_nqtrs	n_per_	_qt z
(a) pelvic_mesh v	0.030	2013-	2017-	18	82	10	92	18	5.1	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.035	2013-	-	18	82	10	92	18	5.1	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.040	-	-	18	81	9	90	18	5.0	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.045	2013-	2017-	18	79	8	87	18	4.8	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.050	2013-	2017-	18	77	8	85	18	4.7	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.055	2013-	2017-	18	77	8	85	18	4.7	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.060	2013-	2017-	18	76	6	82	18	4.6	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.065	2013-	2017-	18	75	5	80	18	4.4	0.1
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.070	2014-	2017-	14	73	3	76	14	5.4	0.0
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.075	2014-	2017-	14	72	3	75	14	5.4	0.0
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.080	2014-	2017-	14	72	3	75	14	5.4	0.0
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.085	2014-	2017-	14	71	2	73	14	5.2	0.0
hernia_mesh		Q3	Q4							
(a) pelvic_mesh v	0.090	2017-	2017-	4	70	1	71	4	17.8	0.0
hernia_mesh		Q1	Q4							
(a) pelvic_mesh v	0.095	2017-	2017-	4	69	1	70	4	17.5	0.0
hernia_mesh		Q1	Q4							
(a) pelvic_mesh v	0.100	2017-	2017-	4	69	1	70	4	17.5	0.0
hernia_mesh		Q1	Q4							
(b) pelvic_mesh v	0.010	2012-	2017-	21	82	59	141	21	6.7	0.7
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.015	2012-	2017-	21	82	59	141	21	6.7	0.7
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.020	2012-	2017-	21	82	56	138	21	6.6	0.7
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.025	2012-	2017-	21	82	56	138	21	6.6	0.7
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.030	2012-	2017-	21	82	55	137	21	6.5	0.7
hernia_mesh/other_mesh		Q4	Q4							

grps	thresh	nmin_	dtmax_	datoews	sum_	_nsAm_	_nt6t_	nqtrs	n_per_	_qt z
(b) pelvic_mesh v	0.035	2012-	2017-	21	82	54	136	21	6.5	0.7
hernia_mesh/other_mesh		Q4	Q4		•	- '				•
(b) pelvic_mesh v	0.040	2012-	2017-	21	81	51	132	21	6.3	0.6
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.045	2012-	2017-	21	79	47	126	21	6.0	0.6
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.050	2012-	2017-	21	77	45	122	21	5.8	0.6
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.055	2013-	2017-	19	77	43	120	19	6.3	0.6
hernia_mesh/other_mesh		Q2	Q4							
(b) pelvic_mesh v	0.060	2013-	2017-	19	76	41	117	19	6.2	0.5
hernia_mesh/other_mesh		Q2	Q4							
(b) pelvic_mesh v	0.065	2013-	2017-	19	75	38	113	19	5.9	0.5
hernia_mesh/other_mesh		Q2	Q4							
(b) pelvic_mesh v	0.070	2013-	2017-	19	73	36	109	19	5.7	0.5
hernia_mesh/other_mesh		Q2	Q4							
(b) pelvic_mesh v	0.075	2013-	2017-	19	72	35	107	19	5.6	0.5
hernia_mesh/other_mesh		Q2	Q4							
(b) pelvic_mesh v	0.080	2014-	2017-	14	72	33	105	14	7.5	0.5
hernia_mesh/other_mesh		Q3	Q4							
(b) pelvic_mesh v	0.085	2014-	2017-	14	71	31	102	14	7.3	0.4
hernia_mesh/other_mesh		Q3	Q4							
(b) pelvic_mesh v	0.090	2014-	2017-	13	70	30	100	13	7.7	0.4
hernia_mesh/other_mesh		Q4	Q4							
(b) pelvic_mesh v	0.095	2014-	2017-	13	69	30	99	13	7.6	0.4
$hernia_mesh/other_mesh$		Q4	Q4							
(b) pelvic_mesh v	0.100	2014-	2017-	13	69	30	99	13	7.6	0.4
$hernia_mesh/other_mesh$		Q4	Q4							
(c) pelvic_mesh v her-	0.010	2012-	2017-	23	82	2017	2099	23	91.3	24.6
$nia_mesh/other_mesh/other_$	device	Q2	Q4							
(c) pelvic_mesh v her-	0.015	2012-	2017-	23	82	1994	2076	5 23	90.3	24.3
$nia_mesh/other_mesh/other_$	device	Q2	Q4							
(c) pelvic_mesh v her-	0.020	2012-	2017-	23	82	1951	2033	3 23	88.4	23.8
$nia_mesh/other_mesh/other_$	device	Q2	Q4							
(c) pelvic_mesh v her-	0.025	2012-	2017-	23	82	1910	1992	2 23	86.6	23.3
$nia_mesh/other_mesh/other_$	device	Q2	Q4							
(c) pelvic_mesh v her-	0.030	2012-	2017-	23	82	1852	1934	1 23	84.1	22.6
$nia_mesh/other_mesh/other_$	device	Q2	Q4							
(c) pelvic_mesh v her-		2012-	2017-	23	82	1783	1865	5 23	81.1	21.7
$nia_mesh/other_mesh/other_$	device	Q2	Q4							

grps	thresh	nmin_o	ltmax_	ditoews	sum_	nsAm_	_nt6tn	nqtrs	n_per_	_qt z
(c) pelvic_mesh v her-	0.040	2012-	2017-	23	81	1715	1796	23	78.1	21.2
nia_mesh/other_mesh/other_d			Q4							
(c) pelvic_mesh v her-	0.045	-	2017-	23	79	1656	1735	23	75.4	21.0
nia_mesh/other_mesh/other_d	levice	Q2	Q4							
(c) pelvic_mesh v her-	0.050	2012-	2017-	23	77	1584	1661	23	72.2	20.6
nia_mesh/other_mesh/other_d	device	Q2	Q4							
(c) pelvic_mesh v her-	0.055	2012-	2017-	23	77	1510	1587	23	69.0	19.6
nia_mesh/other_mesh/other_c	device	Q2	Q4							
(c) pelvic_mesh v her-	0.060	2012-	2017-	23	76	1406	1482	23	64.4	18.5
nia_mesh/other_mesh/other_c	device	Q2	Q4							
(c) pelvic_mesh v her-	0.065	2012-	2017-	23	75	1331	1406	23	61.1	17.7
nia_mesh/other_mesh/other_d	device	Q2	Q4							
(c) pelvic_mesh v her-	0.070	2012-	2017-	23	73	1258	1331	23	57.9	17.2
nia_mesh/other_mesh/other_d	levice	Q2	Q4							
(c) pelvic_mesh v her-	0.075	2012-	2017-	23	72	1186	1258	23	54.7	16.5
nia_mesh/other_mesh/other_d	device	Q2	Q4							
(c) pelvic_mesh v her-	0.080	2012-	2017-	23	72	1104	1176	23	51.1	15.3
nia_mesh/other_mesh/other_d	levice	Q2	Q4							
(c) pelvic_mesh v her-	0.085		2017-	23	71	1024	1095	23	47.6	14.4
nia_mesh/other_mesh/other_d	device	Q2	Q4							
(c) pelvic_mesh v her-	0.090	2012-	2017-	21	70	1013	1083	21	51.6	14.5
nia_mesh/other_mesh/other_d	device	Q4	Q4							
(c) pelvic_mesh v her-	0.095	2012-	2017-	21	69	940	1009	21	48.0	13.6
nia_mesh/other_mesh/other_d	device	Q4	Q4							
(c) pelvic_mesh v her-	0.100	2012-	2017-	21	69	933	1002	21	47.7	13.5
nia_mesh/other_mesh/other_d		Q4	Q4							
(d) hernia_mesh v	0.010	2013-	2017-	20	12	47	59	20	3.0	3.9
other_mesh		Q1	Q4							
(d) hernia_mesh v	0.015	2013-	2017-	20	12	47	59	20	3.0	3.9
other_mesh		Q1	Q4							
(d) hernia_mesh v	0.020	2013-		18	10	46	56	18	3.1	4.6
other_mesh		Q3	Q4							
(d) hernia_mesh v	0.025	2013-	2017-	18	10	46	56	18	3.1	4.6
other_mesh		Q3	Q4							
(d) hernia_mesh v	0.030	2013-	2017-	18	10	45	55	18	3.1	4.5
other_mesh		Q3	Q4							
(d) hernia_mesh v	0.035	2013-	2017-	18	10	44	54	18	3.0	4.4
other_mesh		Q3	Q4							
(d) hernia_mesh v	0.040	2013-	2017-	18	9	42	51	18	2.8	4.7
other_mesh		Q3	Q4							

grps	thresh	min_c	dtmax_	dtews	sum_	nsAm_	_nt6t	nqtrs	n_per_	_qt z
(d) hernia_mesh v	0.045 2	2013-	2017-	18	8	39	47	18	2.6	4.9
other_mesh	(Q3	Q4							
(d) hernia_mesh v	0.050°	2013-	2017-	18	8	37	45	18	2.5	4.6
other_mesh	(Q3	Q4							
(d) hernia_mesh v	0.055 2	2013-	2017-	18	8	35	43	18	2.4	4.4
other_mesh	(Q3	Q4							
(d) hernia_mesh v	0.060°	2013-	2017-	18	6	35	41	18	2.3	5.8
other_mesh	(Q3	Q4							
(d) hernia_mesh v	0.065°	2013-	2017-	18	5	33	38	18	2.1	6.6
other_mesh	(Q3	Q4							
(d) hernia_mesh v	0.070°	2014-	2017-	14	3	33	36	14	2.6	11.0
other_mesh	(Q3	Q4							
(d) hernia_mesh v	0.075 2	2014-	2017-	14	3	32	35	14	2.5	10.
other_mesh	(Q3	Q4							
(d) hernia_mesh v	0.080°	2014-	2017-	13	3	30	33	13	2.5	10.0
other_mesh	(Q4	Q4							
(d) hernia_mesh v	0.085 2	2014-	2017-	13	2	29	31	13	2.4	14.5
other_mesh	(Q4	Q4							
(d) hernia_mesh v	0.090°	2017-	2017-	4	1	29	30	4	7.5	29.0
$other_mesh$	(Q1	Q4							
(d) hernia_mesh v	0.095 2	2017-	2017-	4	1	29	30	4	7.5	29.0
other_mesh	(Q1	Q4							
(d) hernia_mesh v	0.100°	2017-	2017-	4	1	29	30	4	7.5	29.0
other_mesh	(Q1	Q4							
(e) hernia_mesh/other_mesh	0.010°	2012-	2017-	21	59	1958	2017	21	96.0	33.2
v other_device	(Q4	Q4							
(e) hernia_mesh/other_mesh	0.015 2	2012-	2017-	21	59	1935	1994	21	95.0	32.8
v other_device	(Q4	Q4							
(e) hernia_mesh/other_mesh	0.020°	2012-	2017-	21	56	1895	1951	. 21	92.9	33.8
v other_device		Q4	Q4							
(e) hernia_mesh/other_mesh	0.025 2	2012-	2017-	21	56	1854	1910	21	91.0	33.1
v other_device		Q4	Q4							
(e) hernia_mesh/other_mesh	0.030°	2012-	2017-	21	55	1797	1852	21	88.2	32.7
v other_device	(Q4	Q4							
(e) hernia_mesh/other_mesh	0.035 2	2012-	2017-	21	54	1729	1783	21	84.9	32.0
v other_device		Q4	Q4							
(e) hernia_mesh/other_mesh	0.040°	2012-	2017-	21	51	1664	1715	21	81.7	32.6
v other_device	(Q4	Q4							
(e) hernia_mesh/other_mesh	0.045 2	2012-	2017-	21	47	1609	1656	21	78.9	34.2
v other_device	(Q4	Q4							

grps	$threshmin_$	_dtmax	ditews	$\operatorname{sum}_{_}$	_nsAm_	_nt6t	nqtrs	n_per_	_qt z
(e) hernia_mesh/other_mesh	0.050 2012	- 2017-	21	45	1539	1584	21	75.4	34.2
v other_device	Q4	Q4							
(e) hernia_mesh/other_mesh	0.055 2013	- 2017-	19	43	1467	1510	19	79.5	34.1
v other_device	Q2	Q4							
(e) hernia_mesh/other_mesh	$0.060\ 2013$	- 2017-	19	41	1365	1406	19	74.0	33.5
v other_device	Q2	Q4							
(e) hernia_mesh/other_mesh	0.065 2013	- 2017-	19	38	1293	1331	19	70.1	34.0
v other_device	Q2	Q4							
(e) hernia_mesh/other_mesh	0.070 2013	- 2017-	19	36	1222	1258	19	66.2	33.9
v other_device	Q2	Q4							
(e) hernia_mesh/other_mesh	0.075 2013	- 2017-	19	35	1151	1186	19	62.4	32.9
v other_device	Q2	Q4							
(e) hernia_mesh/other_mesh	0.080 2014	- 2017-	14	33	1071	1104	14	78.9	32.5
v other_device	Q3	Q4							
(e) hernia_mesh/other_mesh	0.085 2014	- 2017-	14	31	993	1024	14	73.1	32.0
v other_device	Q3	Q4							
(e) hernia_mesh/other_mesh	$0.090\ 2014$	- 2017-	13	30	983	1013	13	77.9	32.8
v other_device	Q4	Q4							
(e) hernia_mesh/other_mesh	0.095 2014	- 2017-	13	30	910	940	13	72.3	30.3
v other_device	Q4	Q4							
(e) hernia_mesh/other_mesh	$0.100\ 2014$	- 2017-	13	30	903	933	13	71.8	30.1
v other_device	Q4	Q4							

```
# testing/example
row_i <- 1
cv_tab[row_i, ]</pre>
```

```
get_maxsprt_cv(cv_tab$tot_n[row_i], floor(cv_tab$n_per_qtr[row_i]), cv_tab$z[row_i])
```

[1] 3.27782

```
row_i <- 50
  cv_tab[row_i, ]
# A tibble: 1 x 11
            thresh min_dte max_dte rows sum_nA sum_nC tot_n qtrs n_per~1
                                  <int> <dbl> <dbl> <dbl> <int>
  <chr>
            <chr> <chr>
                         <chr>
                                                                      <dbl> <dbl>
1 (c) pelv~ 0.065 2012-Q2 2017-Q4
                                                   1331 1406
                                      23
                                             75
                                                                 23
                                                                       61.1 17.7
# ... with abbreviated variable name 1: n_per_qtr
  get_maxsprt_cv(cv_tab$tot_n[row_i], floor(cv_tab$n_per_qtr[row_i]), cv_tab$z[row_i])
Selected alpha: 0.048 (least conservative value below 0.05)
[1] 2.740269
attr(,"alpha")
[1] 0.048193
  ### takes ~ 70 sec (i5-8400)
  # note purrr::possibly() will just catch when model fails and return as.numeric(NA)
  get_maxsprt_cv_poss <-</pre>
    possibly(get_maxsprt_cv, otherwise = NA_real_, quiet = FALSE)
  tic()
  cv_tab <-
    cv_tab %>%
    # dplyr::filter(row_number() < 7) %>% ### testing
    mutate(
      cv =
        future_pmap_dbl(
          .1 = list(tot_n, floor(n_per_qtr), z),
          .f = \text{~get_maxsprt_cv_poss}(..1, ..2, ..3),
          .options = furrr_seed3
    )
```

Error: For this 'N' there is no solution with prob of Type I error smaller than 0.05. Use 'N $\,$

Error: For this 'N' there is no solution with prob of Type I error smaller than 0.05. Use 'N

```
Error: For this 'N' there is no solution with prob of Type I error smaller than 0.05. Use 'N
Error: For this 'N' there is no solution with prob of Type I error smaller than 0.05. Use 'N
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.047 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.045 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.048 (least conservative value below 0.05)
Selected alpha: 0.046 (least conservative value below 0.05)
```

```
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.045 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.049 (least conservative value below 0.05)
Selected alpha: 0.050 (least conservative value below 0.05)
Selected alpha: 0.048 (least conservative value below 0.05)
```

toc()

176.14 sec elapsed

cv_tab

```
# A tibble: 95 x 12
           thresh min_dte max_dte rows sum_nA sum_nC tot_n qtrs n_per~1
   <chr>
            <chr>
                   <chr>
                           <chr>
                                   <int>
                                          <dbl>
                                                 <dbl> <dbl> <int>
                                                                      <dbl> <dbl>
 1 (a) pel~ 0.010
                   2013-Q1 2017-Q4
                                             82
                                                     12
                                                           94
                                                                       4.7 0.146
                                      20
                                                                 20
2 (a) pel~ 0.015 2013-Q1 2017-Q4
                                      20
                                             82
                                                     12
                                                           94
                                                                 20
                                                                       4.7 0.146
3 (a) pel~ 0.020 2013-Q3 2017-Q4
                                             82
                                                           92
                                                                       5.11 0.122
                                      18
                                                     10
                                                                 18
4 (a) pel~ 0.025 2013-Q3 2017-Q4
                                      18
                                             82
                                                     10
                                                           92
                                                                 18
                                                                       5.11 0.122
5 (a) pel~ 0.030 2013-Q3 2017-Q4
                                      18
                                             82
                                                     10
                                                           92
                                                                 18
                                                                       5.11 0.122
6 (a) pel~ 0.035 2013-Q3 2017-Q4
                                             82
                                                     10
                                      18
                                                           92
                                                                 18
                                                                       5.11 0.122
7 (a) pel~ 0.040 2013-Q3 2017-Q4
                                      18
                                             81
                                                     9
                                                           90
                                                                 18
                                                                       5
                                                                            0.111
8 (a) pel~ 0.045 2013-Q3 2017-Q4
                                             79
                                                     8
                                                                       4.83 0.101
                                      18
                                                           87
                                                                 18
9 (a) pel~ 0.050 2013-Q3 2017-Q4
                                             77
                                                      8
                                                           85
                                                                       4.72 0.104
                                      18
                                                                 18
10 (a) pel~ 0.055 2013-Q3 2017-Q4
                                      18
                                             77
                                                     8
                                                           85
                                                                       4.72 0.104
                                                                 18
# ... with 85 more rows, 1 more variable: cv <dbl>, and abbreviated variable
   name 1: n_per_qtr
  cv_tab %>% dplyr::filter(is.na(cv))
# A tibble: 4 x 12
           thresh min_dte max_dte rows sum_nA sum_nC tot_n qtrs n_per~1
 grps
           <chr> <chr>
                          <chr>
                                  <int>
                                         <dbl>
                                                <dbl> <dbl> <int>
                                                                     <dbl> <dbl>
1 (a) pel~ 0.085 2014-Q3 2017-Q4
                                     14
                                            71
                                                     2
                                                          73
                                                                14
                                                                      5.21 0.0282
                                                                     17.8 0.0143
2 (a) pel~ 0.090 2017-Q1 2017-Q4
                                      4
                                            70
                                                          71
                                                     1
                                                                 4
3 (a) pel~ 0.095 2017-Q1 2017-Q4
                                      4
                                                          70
                                            69
                                                     1
                                                                 4
                                                                     17.5 0.0145
4 (a) pel~ 0.100 2017-Q1 2017-Q4
                                      4
                                            69
                                                     1
                                                          70
                                                                 4
                                                                     17.5 0.0145
# ... with 1 more variable: cv <dbl>, and abbreviated variable name
    1: n_per_qtr
  # remove analyses where thresholds don't allow enough events (extreme threshold values)
  # cv_tab <- cv_tab %>% dplyr::filter(!is.na(cv))
  maxsprt_dat <-
    sra_cum %>%
    mutate(
      maxllr = max\_sprt\_stat\_(c_n = nA, n = nA + nC, z = (nC + nD) / (nA + nB)),
      rre = rr_{est_{(c_n = nA, n = nA + nC, z = (nC + nD) / (nA + nB))}
    )
  # maxsprt_dat
```

```
maxsprt_dat <-
    maxsprt_dat %>%
    left_join(
      ٠,
      cv_tab %>% select(grps, thresh, cv),
      c("grps", "thresh")
    )
  maxsprt_dat <-
    maxsprt_dat %>%
    mutate(
      # some cus don't exist so those llr never reach cu
      reached_cv = if_else(is.na(cv), OL, as.integer(maxllr > cv)),
      # create date for start of each quarter
      dte =
        as_date(paste0(
          substr(mnth, 1, 5),
          sprintf(\%02.0f\%, (as.integer(substr(mnth, 7, 7)) - 1) * 3 + 1),
          "-01"
        ))
    )
  maxsprt_dat %>% dplyr::filter(is.na(cv))
# A tibble: 26 x 13
                                                     nD maxllr
   grps dat_t~1 thresh mnth
                                  nA
                                        nΒ
                                              nC
                                                                 rre
                                                                         cv reach~2
   <chr> <chr>
                 <chr>
                         <chr> <dbl> <dbl> <dbl> <dbl> <
                                                         <dbl> <dbl> <dbl>
                                                                              <int>
1 (a) ~ cumula~ 0.085
                        2014~
                                        20
                                                     23
                                                          1.79 5.54
                                                                                  0
                                   6
                                                1
                                                                         NA
2 (a) ~ cumula~ 0.085
                                                                                  0
                        2014~
                                  23
                                        22
                                                     30
                                                          8.79 15.8
                                                                         NA
3 (a) ~ cumula~ 0.085
                        2015~
                                  24
                                        22
                                                     31
                                                          9.37 16.7
                                                                                  0
                                                                         NA
4 (a) ~ cumula~ 0.085
                        2015~
                                  24
                                        22
                                                     31
                                                          9.37 16.7
                                                                         NA
                                                                                  0
5 (a) ~ cumula~ 0.085
                        2015~
                                  24
                                        23
                                                1
                                                     31
                                                          9.17 16.3
                                                                         NA
                                                                                  0
6 (a) ~ cumula~ 0.085
                        2015~
                                        23
                                                          9.17 16.3
                                                                                  0
                                  24
                                                1
                                                     31
                                                                         NΑ
7 (a) ~ cumula~ 0.085
                        2016~
                                  31
                                        24
                                                1
                                                     33 11.4 19.2
                                                                         NA
                                                                                  0
8 (a) ~ cumula~ 0.085
                                                1
                                                     33 11.4 19.2
                        2016~
                                  31
                                        24
                                                                         NA
                                                                                  0
9 (a) ~ cumula~ 0.085
                        2016~
                                  33
                                        26
                                                1
                                                     39
                                                         13.5 22.4
                                                                         NA
                                                                                  0
10 (a) ~ cumula~ 0.085 2016~
                                                         13.5 22.4
                                  33
                                        26
                                                1
                                                     39
                                                                                  0
                                                                         NA
# ... with 16 more rows, 1 more variable: dte <date>, and abbreviated variable
    names 1: dat_type, 2: reached_cv
```

```
# have a peak
  maxsprt_dat %>%
    select(-dat_type) %>%
    print(., n = 25)
# A tibble: 1,707 x 12
                                                 nC
                                                        nD maxllr
   grps
                  thresh mnth
                                    nA
                                           nB
                                                                             cv reach~1
                                                                     rre
   <chr>
                  <chr>
                          <chr> <dbl> <dbl> <dbl> <dbl> <dbl>
                                                            <dbl> <dbl> <dbl>
                                                                                   <int>
 1 (a) pelvic m~ 0.010
                          2013~
                                            7
                                                  1
                                                         4 0.0657
                                                                           3.28
                                                                                       0
                                     3
                                                                    1.5
 2 (a) pelvic_m~ 0.010
                          2013~
                                     4
                                           10
                                                  1
                                                         5 0.129
                                                                    1.71
                                                                           3.28
                                                                                       0
                                                  2
                                                         9 0.224
3 (a) pelvic_m~ 0.010
                          2013~
                                     5
                                           11
                                                                    1.72
                                                                           3.28
                                                                                       0
4 (a) pelvic_m~ 0.010
                          2013~
                                     9
                                           11
                                                  2
                                                         9 0.801
                                                                    2.48
                                                                           3.28
                                                                                       0
5 (a) pelvic_m~ 0.010
                                     9
                                                  2
                                                        10 0.976
                                                                           3.28
                          2014~
                                           11
                                                                    2.7
                                                                                       0
6 (a) pelvic_m~ 0.010
                          2014~
                                    10
                                           12
                                                  3
                                                        12 0.885
                                                                    2.27
                                                                           3.28
                                                                                       0
7 (a) pelvic_m~ 0.010
                          2014~
                                    12
                                           14
                                                  5
                                                        19 1.22
                                                                    2.22
                                                                           3.28
                                                                                       0
                                                  7
8 (a) pelvic_m~ 0.010
                                                        24 4.05
                                                                    2.95
                          2014~
                                    30
                                           15
                                                                           3.28
                                                                                       1
                                                  7
9 (a) pelvic_m~ 0.010
                          2015~
                                                        25 4.45
                                                                           3.28
                                    31
                                           15
                                                                    3.08
                                                                                       1
10 (a) pelvic_m~ 0.010
                          2015~
                                    31
                                           15
                                                  7
                                                        25 4.45
                                                                    3.08
                                                                           3.28
                                                                                       1
                                                  7
11 (a) pelvic_m~ 0.010
                          2015~
                                    31
                                           16
                                                        25 4.27
                                                                    3.02
                                                                           3.28
                                                                                       1
12 (a) pelvic_m~ 0.010
                          2015~
                                                  9
                                                        25 3.36
                                                                    2.49
                                                                           3.28
                                    31
                                           16
                                                                                       1
13 (a) pelvic_m~ 0.010
                          2016~
                                    35
                                           16
                                                  9
                                                        25 3.83
                                                                    2.59
                                                                           3.28
                                                                                       1
14 (a) pelvic m~ 0.010
                          2016~
                                    39
                                                  9
                                                        25 4.27
                                           16
                                                                    2.68
                                                                           3.28
                                                                                       1
15 (a) pelvic_m~ 0.010
                          2016~
                                    39
                                           16
                                                 10
                                                        30 5.17
                                                                    2.84
                                                                           3.28
                                                                                       1
16 (a) pelvic m~ 0.010
                          2016~
                                                        30 5.04
                                                                    2.78
                                                                           3.28
                                                                                       1
                                    41
                                           18
                                                 10
17 (a) pelvic_m~ 0.010
                          2017~
                                    50
                                           18
                                                 11
                                                        31 5.85
                                                                    2.81
                                                                           3.28
                                                                                       1
18 (a) pelvic_m~ 0.010
                          2017~
                                    63
                                           19
                                                 12
                                                        33 7.04
                                                                    2.88
                                                                           3.28
                                                                                       1
19 (a) pelvic_m~ 0.010
                          2017~
                                    73
                                           19
                                                 12
                                                        34 8.18
                                                                    3.04
                                                                           3.28
                                                                                       1
                                          20
20 (a) pelvic_m~ 0.010
                          2017~
                                    82
                                                 12
                                                        34 8.65
                                                                    3.08
                                                                           3.28
                                                                                       1
21 (a) pelvic_m~ 0.015
                          2013~
                                     3
                                            7
                                                  1
                                                         4 0.0657
                                                                    1.5
                                                                           3.28
                                                                                       0
22 (a) pelvic_m~ 0.015
                          2013~
                                     4
                                           10
                                                  1
                                                         5 0.129
                                                                           3.28
                                                                                       0
                                                                    1.71
23 (a) pelvic_m~ 0.015
                          2013~
                                     5
                                                  2
                                                         9 0.224
                                                                    1.72
                                                                           3.28
                                                                                       0
                                           11
                                                  2
24 (a) pelvic_m~ 0.015
                          2013~
                                     9
                                                         9 0.801
                                                                    2.48
                                                                           3.28
                                                                                       0
                                           11
25 (a) pelvic_m~ 0.015
                          2014~
                                     9
                                                  2
                                                        10 0.976
                                           11
                                                                    2.7
                                                                           3.28
                                                                                       0
# ... with 1,682 more rows, 1 more variable: dte <date>, and abbreviated
    variable name 1: reached_cv
  # first signif
  maxsprt_signif <-</pre>
    maxsprt_dat %>%
```

```
group_by(grps, dat_type, thresh) %>%
dplyr::filter(reached_cv > 0) %>%
```

```
arrange(dte) %>%
    dplyr::filter(row_number() == 1) %>%
    ungroup() %>%
    rename(dte_reach_sig = dte)
  nrow(maxsprt dat)
[1] 1707
  maxsprt_dat <-
    left_join(
      maxsprt_dat,
      maxsprt_signif %>% select(grps, dat_type, thresh, dte_reach_sig),
      c("grps", "dat_type", "thresh")
    )
  nrow(maxsprt_dat)
[1] 1707
  maxsprt_dat
# A tibble: 1,707 x 14
   grps dat_t~1 thresh mnth
                                        nB
                                              nC
                                                     nD maxllr
                                                                         cv reach~2
                                  nA
                                                                  rre
   <chr> <chr>
                 <chr>>
                         <chr> <dbl> <dbl> <dbl> <dbl> <dbl>
                                                         <dbl> <dbl> <dbl>
                                                                              <int>
 1 (a) ~ cumula~ 0.010
                        2013~
                                   3
                                         7
                                                1
                                                      4 0.0657
                                                                1.5
                                                                       3.28
                                                                                  0
2 (a) ~ cumula~ 0.010
                        2013~
                                        10
                                                1
                                                      5 0.129
                                                                1.71
                                                                       3.28
                                                                                  0
                                   4
3 (a) ~ cumula~ 0.010 2013~
                                   5
                                        11
                                                2
                                                      9 0.224
                                                                1.72
                                                                       3.28
                                                                                  0
4 (a) ~ cumula~ 0.010 2013~
                                   9
                                        11
                                                2
                                                      9 0.801
                                                                2.48
                                                                       3.28
                                                                                  0
5 (a) ~ cumula~ 0.010 2014~
                                   9
                                                2
                                                     10 0.976
                                                                       3.28
                                                                                  0
                                        11
                                                                2.7
6 (a) ~ cumula~ 0.010 2014~
                                  10
                                        12
                                                3
                                                     12 0.885
                                                                2.27
                                                                       3.28
                                                                                  0
7 (a) ~ cumula~ 0.010 2014~
                                                5
                                                     19 1.22
                                                                2.22
                                                                                  0
                                  12
                                        14
                                                                       3.28
8 (a) ~ cumula~ 0.010 2014~
                                                7
                                                     24 4.05
                                                                2.95
                                                                       3.28
                                                                                  1
                                  30
                                        15
9 (a) ~ cumula~ 0.010
                                                7
                        2015~
                                  31
                                        15
                                                     25 4.45
                                                                3.08
                                                                      3.28
                                                                                  1
10 (a) ~ cumula~ 0.010 2015~
                                  31
                                        15
                                                7
                                                     25 4.45
                                                                3.08
                                                                      3.28
                                                                                  1
# ... with 1,697 more rows, 2 more variables: dte <date>, dte_reach_sig <date>,
    and abbreviated variable names 1: dat_type, 2: reached_cv
```

```
maxsprt dat <-
    maxsprt_dat %>%
    mutate(
      dte_reach_sig = if_else(is.na(dte_reach_sig), as_date(today()), dte_reach_sig),
      reach_sig = dte >= dte_reach_sig
    )
  # these are where the maxllr has dropped under the CV after exceeding it previously
  maxsprt_dat %>%
    dplyr::filter(
      is.na(reach_sig) |
        is.na(reached cv) |
        (as.logical(reached_cv) != reach_sig)
    )
# A tibble: 5 x 15
  grps
        dat_t~1 thresh mnth
                                 nA
                                       nΒ
                                             nC
                                                   nD maxllr
                                                               rre
                <chr> <chr> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <
1 (c) p~ cumula~ 0.085 2013~
                                        7
                                             84 1449
                                                        2.61 5.48
                                                                    2.88
                                                                               0
                                  3
2 (c) p~ cumula~ 0.085 2013~
                                        7
                                                                               0
                                  3
                                             84 1449
                                                        2.61 5.48
                                                                    2.88
3 (c) p~ cumula~ 0.085 2013~
                                  3
                                       7
                                             84 1449
                                                        2.61 5.48
                                                                   2.88
                                                                               0
4 (c) p~ cumula~ 0.085 2013~
                                  3
                                        7
                                             84 1449
                                                        2.61 5.48 2.88
                                                                               0
                                            772 7436
                                                        2.71 2.01 2.71
                                                                               0
5 (e) h~ cumula~ 0.065 2015~
                                 14
                                       60
# ... with 3 more variables: dte <date>, dte_reach_sig <date>, reach_sig <lgl>,
    and abbreviated variable names 1: dat_type, 2: reached_cv
  maxsprt_dat <-
    maxsprt_dat %>%
    select(-reached_cv)
  maxsprt_dat %>%
    write_parquet(., sink = "out/sra_cum_maxsprt.parquet")
```

3 Ready plot data

```
plt_dat <-
 bind_rows(
    maxsprt_dat %>%
      select(comparator, dte, cv, reached_cv, val = maxllr) %>%
      mutate(stat = "MaxSPRT (max LLR)"),
    bcpnn data %>%
      select(comparator, dte, val = ci_lo) %>%
      mutate(cv = 0, reached_cv = as.integer(val > cv), stat = "IC (BCPNN, Lower 95% CI)")
  )
sig_reach_dat <-
 plt_dat %>%
  arrange(stat, comparator, dte) %>%
  group_by(stat, comparator) %>%
  dplyr::filter(reached_cv == 1) %>%
  dplyr::filter(row_number() == 1) %>%
  select(stat, comparator, dte_reached = dte) %>%
  # now create separation between reached CV values when it occurs
  group_by(stat, dte_reached) %>%
 mutate(rep_dte = 1:n()) %>%
  ungroup() %>%
  mutate(dte_reached = dte_reached + days(10 * (rep_dte - 1))) %>%
  select(-rep_dte)
plt_dat <-
 left_join(
   plt_dat,
    sig_reach_dat,
    c("stat", "comparator")
plt_dat %>%
  ggplot(., aes(x = dte, y = val, col = comparator)) +
  geom_hline(aes(yintercept = cv), alpha = 0.5) +
  geom_vline(aes(xintercept = dte_reached, col = comparator), alpha = 0.5) +
  geom_line(alpha = 0.5) +
  geom_point() +
```

```
facet_wrap(~ stat, ncol = 1, scales = "free_y") +
scale_colour_tableau() +
theme_bw() +
labs(
    x = "Quarter",
    y = "Statistic",
    col = "Comparison"
)
```

4 Session information

```
# Sys.info()[!(names(Sys.info()) %in% c("login", "nodename"))] %>%
  # as.data.frame(.)
  format(Sys.time(), '%d %b %Y')
[1] "05 Nov 2023"
  sessionInfo()
R version 4.2.2 (2022-10-31 ucrt)
Platform: x86_64-w64-mingw32/x64 (64-bit)
Running under: Windows 10 x64 (build 19045)
Matrix products: default
locale:
[1] LC_COLLATE=English_Australia.utf8 LC_CTYPE=English_Australia.utf8
[3] LC_MONETARY=English_Australia.utf8 LC_NUMERIC=C
[5] LC_TIME=English_Australia.utf8
attached base packages:
[1] stats
              graphics grDevices utils
                                            datasets methods
                                                                 base
other attached packages:
 [1] pharmsignal_0.1.0 arrow_11.0.0.2
                                         foreach_1.5.2
                                                            gsDesign_3.5.0
 [5] knitr_1.42
                       ggrepel_0.9.3
                                         ggplot2_3.4.1
                                                            tictoc_1.2
 [9] lubridate_1.9.2 furrr_0.3.1
                                         future 1.33.0
                                                            purrr 1.0.1
[13] forcats_1.0.0
                      tidyr_1.3.0
                                         dplyr_1.1.0
                                                            readr_2.1.4
loaded via a namespace (and not attached):
 [1] Rcpp_1.0.10
                                lattice_0.20-45
 [3] listenv_0.9.0
                                assertthat_0.2.1
 [5] digest_0.6.31
                                utf8_1.2.3
 [7] parallelly_1.36.0
                                R6_2.5.1
 [9] MatrixModels_0.5-1
                                evaluate_0.20
[11] coda_0.19-4
                                pillar_1.8.1
[13] rlang_1.0.6
                                rstudioapi_0.14
[15] SparseM_1.81
                                Matrix_1.5-1
[17] rmarkdown_2.20
                                splines_4.2.2
```

[19] bit_4.0.5 munsell_0.5.0 [21] compiler_4.2.2 xfun_0.37 [23] pkgconfig_2.0.3 EmpiricalCalibration_3.1.1 [25] globals_0.16.2 $mcmc_0.9-7$ [27] htmltools_0.5.4 tidyselect_1.2.0 [29] tibble_3.1.8 codetools_0.2-18 [31] fansi_1.0.4 tzdb_0.3.0 [33] withr_2.5.0 MASS_7.3-58.1 [35] grid_4.2.2 jsonlite_1.8.4 [37] xtable_1.8-4 gtable_0.3.1 [39] lifecycle_1.0.3 magrittr_2.0.3 [41] scales_1.2.1 cli_3.6.0 [43] xml2_1.3.3 ellipsis_0.3.2 [45] generics_0.1.3 vctrs_0.5.2 [47] boot_1.3-28 iterators_1.0.14 [49] tools_4.2.2 bit64_4.0.5 [51] glue_1.6.2 $hms_1.1.2$ [53] parallel_4.2.2 fastmap_1.1.0 [55] survival_3.4-0 yam1_2.3.7 [57] timechange_0.2.0 colorspace_2.1-0

gt_0.10.0

MCMCpack_1.6-3

[59] Sequential_4.3.1

[61] quantreg_5.94