Data analysis

Signal detection of spontaneous medical device reports over time

Ty Stanford et al.

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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 'future' 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")
```

1.2 Parallel compution setup

```
# options(future.globals.maxSize = 500 * 1024 ^ 2) # = 500 MiB
options(future.globals.maxSize = 1e3 * 1024 ^ 2) # = 1 GB
```

```
# furrr parallel workers/cores setup
# change `workers = 4` based on cores available in processor being used
plan(multisession, workers = 4)

### 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>
```

6.12 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</pre>
```

2.94 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)</pre>
```

1.3 Constants

```
# arbitrarily, let's go with minimum cell count of 3 (should be discussed!)
arbitrary_cell_min <- 1</pre>
```

1.4 Functions

```
get_sig_tab <- function(nA, nB, nC, nD, alpha = 0.05, n_mcmc = 1e+05) {</pre>
  out_cols_of_interest <- c("est_name", "est_scale", "est", "ci_lo", "ci_hi")</pre>
  sig_tab <- pharmsignal::bcpnn_mcmc_signal(nA, nB, nC, nD, alpha = alpha, n_mcmc = n_mcmc</pre>
  sig_tab <- sig_tab[, out_cols_of_interest]</pre>
  # sig_tab <- bind_cols(tibble(mnth = mnth), sig_tab)</pre>
  return(sig_tab)
}
get_sig_tab_over_time <- function(dat, alpha = 0.05, 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 = alpha, n_mcmc = n_mcmc
        )
      )
    }
  return(sig_tab_over_time)
}
```

1.5 Load data

```
sra_dat <- read_parquet("dat/sra_dat.parquet")</pre>
```

2 Analysis

2.1 BCPNN

```
sra_cum <-
    sra_dat %>%
    dplyr::filter(dat_type == "cumulative")
  # make data for each combination of params nested for purrr like processing
  sra_cum <-
    sra_cum %>%
    nest(data = c(mnth, nA, nB, nC, nD))
  # testing/example
  sra_cum$data[[1]]
# A tibble: 44 x 5
  mnth
              nA
                    nΒ
                          nC
                                 nD
   <chr>
           <dbl> <dbl> <dbl> <dbl> <dbl>
 1 2013-01
               3
                     7
                                  2
                            1
                     7
2 2013-02
               3
                            1
                                  4
                     7
                                  5
3 2013-04
               3
4 2013-05
                    10
                                  5
                                  7
5 2013-07
                    11
                           1
6 2013-08
               5
                                  7
                    11
                           1
7 2013-09
               5
                    11
                            2
                                  9
8 2013-11
               8
                    11
                            2
                                  9
9 2013-12
               9
                    11
                            2
                                  9
10 2014-03
                                 10
                    11
# ... with 34 more rows
  get_sig_tab_over_time(sra_cum$data[[1]])
     est_name est_scale
                                 est
                                            ci_lo
                                                       ci_hi
1 bcpnn_mcmc
                   log2 -0.03155473 -1.008567292 0.5011816
                   log2 0.14423111 -0.881461274 0.7657139
2 bcpnn_mcmc
3 bcpnn_mcmc
                   log2 0.22225892 -0.819459061 0.8710605
4 bcpnn_mcmc
                   log2 0.16982567 -0.631979264 0.6843789
                   log2 0.25877437 -0.558250981 0.7901982
5 bcpnn_mcmc
```

```
0.28945108 -0.359347606 0.7778787
6
   bcpnn_mcmc
                    log2
7
   bcpnn_mcmc
                    log2
                          0.24275055 -0.534420549 0.7826432
                          0.31486271 -0.170553281 0.7465905
8
   bcpnn_mcmc
                   log2
                          0.32253264 -0.103384822 0.7293047
9
   bcpnn_mcmc
                    log2
10 bcpnn mcmc
                    log2
                          0.36527691 -0.067632484 0.7879723
11 bcpnn_mcmc
                    log2
                          0.40661823 -0.034175482 0.8369273
12 bcpnn_mcmc
                    log2
                          0.31431083 -0.117133215 0.7114928
13 bcpnn_mcmc
                    log2
                          0.35146856 -0.088210721 0.7597945
14 bcpnn_mcmc
                    log2
                          0.32236631 -0.145358061 0.7374164
15 bcpnn_mcmc
                    log2
                          0.41872427
                                      0.005768877 0.8060994
                          0.42043314 -0.019393480 0.8193463
16 bcpnn_mcmc
                    log2
  bcpnn_mcmc
                    log2
                          0.51134622
                                       0.122251507 0.8922410
                          0.48997255
                                       0.090959174 0.8764148
18 bcpnn_mcmc
                    log2
19 bcpnn_mcmc
                    log2
                          0.44480211
                                       0.225394906 0.6963962
20 bcpnn_mcmc
                    log2
                          0.46323762
                                       0.241125813 0.7192630
21 bcpnn_mcmc
                    log2
                          0.45935820
                                       0.241366445 0.7109985
                    log2
                          0.44698469
                                       0.231636145 0.6971812
22 bcpnn_mcmc
                                       0.211373941 0.6762936
23 bcpnn_mcmc
                    log2
                          0.42807282
                                       0.191379522 0.6565181
24 bcpnn_mcmc
                    log2
                          0.40986367
                          0.40529856
                                       0.198145687 0.6409685
25 bcpnn mcmc
                    log2
26 bcpnn_mcmc
                    log2
                          0.40021287
                                       0.201086693 0.6289638
27 bcpnn mcmc
                    log2
                          0.38907653
                                       0.206723284 0.6022235
28 bcpnn_mcmc
                    log2
                          0.42060041
                                       0.232653673 0.6397847
29 bcpnn_mcmc
                    log2
                          0.43735676
                                       0.243979517 0.6600397
                          0.45236046
                                       0.256382608 0.6771721
30 bcpnn_mcmc
                    log2
                    log2
                                       0.249050440 0.6506842
31 bcpnn_mcmc
                          0.43564014
32 bcpnn_mcmc
                    log2
                          0.42578750
                                       0.240710552 0.6406515
33 bcpnn_mcmc
                    log2
                          0.41437069
                                       0.240937476 0.6170948
34 bcpnn_mcmc
                    log2
                          0.40562061
                                       0.238810116 0.5993361
                          0.40236748
                                       0.238325445 0.5936070
35 bcpnn_mcmc
                    log2
36 bcpnn_mcmc
                    log2
                          0.40588775
                                       0.250241656 0.5864129
                          0.38625000
                                       0.238767885 0.5591967
37 bcpnn_mcmc
                    log2
38 bcpnn_mcmc
                    log2
                          0.37617897
                                       0.235058228 0.5426621
                                       0.243192323 0.5467141
39 bcpnn_mcmc
                    log2
                          0.38226092
                                       0.235354142 0.5233362
40 bcpnn mcmc
                    log2
                          0.36736813
41 bcpnn_mcmc
                   log2
                          0.36257011
                                       0.233573242 0.5155766
42 bcpnn mcmc
                    log2
                          0.35553674
                                       0.229417693 0.5041057
43 bcpnn_mcmc
                          0.34192103
                                       0.220789503 0.4841618
                    log2
44 bcpnn_mcmc
                    log2
                          0.33766947
                                       0.219170482 0.4782721
```

112.24 sec elapsed

```
# check
sra_cum$sig_tab[[1]]
```

```
est_name est_scale
                                est
                                          ci_lo
                                                    ci_hi
                   log2 -0.03155473 -1.01361518 0.5008338
1 bcpnn_mcmc
2 bcpnn_mcmc
                         0.14423111 -0.87754425 0.7674485
                   log2
  bcpnn_mcmc
                   log2
                         0.22225892 -0.82925568 0.8768976
3
                         0.16982567 -0.62630109 0.6841108
  bcpnn_mcmc
                   log2
  bcpnn_mcmc
                   log2
                         0.25877437 -0.56115069 0.7903408
                         0.28945108 -0.36195188 0.7791041
6
  bcpnn_mcmc
                   log2
7
  bcpnn_mcmc
                   log2 0.24275055 -0.54147551 0.7865138
                   log2 0.31486271 -0.16358209 0.7464793
8
  bcpnn_mcmc
  bcpnn_mcmc
                   log2
                         0.32253264 -0.09929240 0.7367417
10 bcpnn_mcmc
                   log2
                         0.36527691 -0.06997405 0.7826290
                   log2 0.40661823 -0.03559667 0.8371538
11 bcpnn_mcmc
12 bcpnn_mcmc
                   log2
                         0.31431083 -0.12150818 0.7158149
13 bcpnn_mcmc
                   log2
                         0.35146856 -0.08877886 0.7628745
14 bcpnn mcmc
                   log2 0.32236631 -0.15065287 0.7402075
15 bcpnn_mcmc
                   log2 0.41872427 0.00681964 0.8113143
                         0.42043314 -0.01475558 0.8214752
16 bcpnn_mcmc
                   log2
17 bcpnn_mcmc
                   log2 0.51134622 0.12474650 0.8952461
                   log2 0.48997255 0.09205215 0.8748550
18 bcpnn_mcmc
19 bcpnn_mcmc
                                     0.22642533 0.6997523
                   log2
                         0.44480211
20 bcpnn_mcmc
                   log2
                         0.46323762  0.24023405  0.7193275
```

```
21 bcpnn_mcmc
                   log2
                         0.45935820
                                     0.24200791 0.7127425
22 bcpnn_mcmc
                   log2
                         0.44698469
                                     0.23167170 0.6944536
                         0.42807282
                                     0.21226736 0.6772533
23 bcpnn_mcmc
                   log2
                                     0.19249103 0.6557647
24 bcpnn_mcmc
                   log2
                         0.40986367
                   log2
25 bcpnn mcmc
                         0.40529856
                                     0.19777767 0.6411809
26 bcpnn_mcmc
                                     0.20072109 0.6295022
                   log2
                         0.40021287
27 bcpnn_mcmc
                   log2
                         0.38907653
                                     0.20484737 0.6038758
28 bcpnn_mcmc
                   log2
                         0.42060041
                                     0.23274148 0.6415478
29 bcpnn_mcmc
                   log2
                         0.43735676
                                     0.24612622 0.6588602
30 bcpnn_mcmc
                   log2
                         0.45236046
                                     0.25818115 0.6760420
31 bcpnn_mcmc
                   log2
                         0.43564014
                                     0.24895062 0.6536131
                                     0.23965009 0.6400406
32 bcpnn_mcmc
                   log2
                         0.42578750
                         0.41437069
                                     0.23951271 0.6165066
33 bcpnn_mcmc
                   log2
34 bcpnn_mcmc
                   log2
                         0.40562061
                                     0.23923085 0.5994463
35 bcpnn_mcmc
                   log2
                         0.40236748
                                     0.23938575 0.5932197
                         0.40588775
                                     0.24969119 0.5872826
36 bcpnn_mcmc
                   log2
37 bcpnn_mcmc
                   log2
                         0.38625000
                                     0.23831821 0.5599170
                         0.37617897
                                     0.23431857 0.5426523
38 bcpnn_mcmc
                   log2
39 bcpnn_mcmc
                         0.38226092
                                     0.24172534 0.5480638
                   log2
40 bcpnn mcmc
                         0.36736813
                                     0.23617735 0.5236761
                   log2
41 bcpnn_mcmc
                   log2
                         0.36257011
                                     0.23406734 0.5142182
42 bcpnn_mcmc
                   log2
                         0.35553674
                                     0.22958272 0.5048607
43 bcpnn_mcmc
                   log2
                         0.34192103
                                     0.22109856 0.4837613
44 bcpnn_mcmc
                                     0.22005543 0.4760598
                   log2
                         0.33766947
  sra_cum_bcpnn <-</pre>
    sra_cum %>%
    unnest(cols = c(data, sig_tab)) %>%
    mutate(dte = as date(paste0(mnth, "-01")))
  sra_cum_bcpnn
```

A tibble: 4,523 x 14

	grps	dat_t~1	thresh	mnth	nA	nB	nC	nD	est_n~2	est_s~3	est
	<chr></chr>	<chr></chr>	<dbl></dbl>	<chr>></chr>	<dbl></dbl>	<dbl></dbl>	<dbl></dbl>	<dbl></dbl>	<chr></chr>	<chr></chr>	<dbl></dbl>
1	pelvic_~	cumula~	0.01	2013~	3	7	1	2	bcpnn_~	log2	-0.0316
2	pelvic_~	cumula~	0.01	2013~	3	7	1	4	bcpnn_~	log2	0.144
3	pelvic_~	cumula~	0.01	2013~	3	7	1	5	bcpnn_~	log2	0.222
4	pelvic_~	cumula~	0.01	2013~	4	10	1	5	bcpnn_~	log2	0.170
5	pelvic_~	cumula~	0.01	2013~	4	11	1	7	bcpnn_~	log2	0.259
6	pelvic_~	cumula~	0.01	2013~	5	11	1	7	bcpnn_~	log2	0.289

```
5
7 pelvic_~ cumula~
                      0.01 2013~
                                           11
                                                  2
                                                        9 bcpnn_~ log2
                                                                            0.243
8 pelvic_~ cumula~
                      0.01 2013~
                                           11
                                                  2
                                                        9 bcpnn_~ log2
                                                                            0.315
                                      8
9 pelvic_~ cumula~
                      0.01 2013~
                                                  2
                                                        9 bcpnn_~ log2
                                      9
                                           11
                                                                            0.323
10 pelvic_~ cumula~
                      0.01 2014~
                                      9
                                           11
                                                  2
                                                       10 bcpnn_~ log2
                                                                            0.365
# ... with 4,513 more rows, 3 more variables: ci_lo <dbl>, ci_hi <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] 4523
  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] 4523
  sra_cum_bcpnn
# A tibble: 4,523 x 15
            dat_t~1 thresh mnth
                                                 nC
                                   \mathtt{n}\mathtt{A} \mathtt{n}\mathtt{B}
                                                       nD est_n~2 est_s~3
  grps
                                                                               est
```

```
<dbl> <chr> <dbl> <dbl> <dbl> <dbl> <dbl> <chr>
   <chr>
            <chr>
                                                                           <dbl>
 1 pelvic_~ cumula~
                      0.01 2013~
                                           7
                                     3
                                                 1
                                                       2 bcpnn_~ log2
                                                                         -0.0316
                                           7
2 pelvic_~ cumula~
                      0.01 2013~
                                     3
                                                 1
                                                       4 bcpnn_~ log2
                                                                          0.144
3 pelvic_~ cumula~
                      0.01 2013~
                                           7
                                                 1
                                                       5 bcpnn_~ log2
                                                                          0.222
                                     3
4 pelvic ~ cumula~
                                                 1
                                                       5 bcpnn ~ log2
                      0.01 2013~
                                     4
                                          10
                                                                          0.170
5 pelvic_~ cumula~
                      0.01 2013~
                                                 1
                                                       7 bcpnn_~ log2
                                                                          0.259
                                     4
                                          11
                                                     7 bcpnn_~ log2
6 pelvic ~ cumula~
                      0.01 2013~
                                     5
                                          11
                                                 1
                                                                          0.289
7 pelvic_~ cumula~
                                                       9 bcpnn_~ log2
                      0.01 2013~
                                     5
                                          11
                                                 2
                                                                          0.243
8 pelvic_~ cumula~
                      0.01 2013~
                                          11
                                                 2
                                                       9 bcpnn_~ log2
                                                                          0.315
                                     8
9 pelvic_~ cumula~
                                                 2
                                                       9 bcpnn_~ log2
                      0.01 2013~
                                     9
                                          11
                                                                          0.323
10 pelvic_~ cumula~
                      0.01 2014~
                                                 2
                                                      10 bcpnn_~ log2
                                                                          0.365
                                     9
                                          11
# ... with 4,513 more rows, 4 more variables: ci_lo <dbl>, ci_hi <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
    )
```

2.2 BCPNN with mult comp adjust

2.3 MaxSPRT

3 Session information

```
sra_cum_bcpnn %>%
    write_parquet(., sink = "out/sra_cum_bcpnn.parquet")
  ## close multisession workers by switching plan
  plan(sequential)
  format(Sys.time(), '%d %b %Y')
[1] "19 Jun 2023"
  Sys.info() %>% as.data.frame(.)
                       Windows
sysname
release
                       10 x64
version
                   build 19044
nodename
             DESKTOP-R5P5N23
machine
                        x86-64
login
                            ty
user
                            ty
effective_user
                            ty
  sessionInfo()
R version 4.2.2 (2022-10-31 ucrt)
Platform: x86_64-w64-mingw32/x64 (64-bit)
Running under: Windows 10 x64 (build 19044)
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_3.4.0
[5]	knitr_1.42	ggrepel_0.9.3	ggplot2_3.4.1	tictoc_1.1
[9]	<pre>lubridate_1.9.2</pre>	furrr_0.3.1	future_1.32.0	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]	tidyselect_1.2.0	xfun_0.37	listenv_0.9.0	splines_4.2.2
[5]	lattice_0.20-45	colorspace_2.1-0	vctrs_0.5.2	<pre>generics_0.1.3</pre>
[9]	htmltools_0.5.4	yam1_2.3.7	survival_3.4-0	MCMCpack_1.6-3
[13]	utf8_1.2.3	rlang_1.0.6	pillar_1.8.1	glue_1.6.2
[17]	withr_2.5.0	bit64_4.0.5	lifecycle_1.0.3	MatrixModels_0.5-1
[21]	munsell_0.5.0	gtable_0.3.1	coda_0.19-4	codetools_0.2-18
[25]	evaluate_0.20	SparseM_1.81	tzdb_0.3.0	fastmap_1.1.0
[29]	quantreg_5.94	parallel_4.2.2	fansi_1.0.4	Rcpp_1.0.10
[33]	xtable_1.8-4	scales_1.2.1	jsonlite_1.8.4	parallelly_1.34.0
[37]	bit_4.0.5	$mcmc_0.9-7$	hms_1.1.2	digest_0.6.31
[41]	grid_4.2.2	cli_3.6.0	tools_4.2.2	magrittr_2.0.3
[45]	tibble_3.1.8	pkgconfig_2.0.3	Matrix_1.5-3	MASS_7.3-58.1
[49]	ellipsis_0.3.2	timechange_0.2.0	$assertthat_0.2.1$	rmarkdown_2.20
[53]	rstudioapi_0.14	iterators_1.0.14	R6_2.5.1	globals_0.16.2
[57]	compiler_4.2.2			