# Transporter Colocalization Analysis (ImageJ Coloc2)

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### 2022-12-01

M1 = occludin overlapping P-gp/Bcrp/Mrp2 M2 = P-gp/Bcrp/Mrp2 overlapping occludin

# Reading in the data

# P-gp

```
#starting with the raw data
#pqp_filelist <- list.files(path="../data/IHC/coloc2_results_pqp/",</pre>
                            pattern="*.txt", full.names = TRUE)
#pgp_cols <- c("sample", "m1", "m2", "r_pearson", "Pval", "r_rand", "sd_rand")
#pgp_coloc2 <- as.data.frame(matrix(,1,length(pgp_cols)))</pre>
#names(pgp_coloc2) <- pgp_cols</pre>
#for (i in pgp_filelist) {
# test <- read lines(i)</pre>
# sample <- str_subset(test, "Working on") %>%
   str_extract(., "(?<=:[:space:]).*")
# r_pearson <- str_subset(test, "no threshold") %>%
   str extract(., "(?<=,[:space:]).*")
# m1 <- str_subset(test, "tM1") %>% str_extract(., "(?<=,[:space:]).*")
# m2 <- str_subset(test, "tM2") %>% str_extract(., "(?<=,[:space:]).*")
\# Pval <- str_subset(test, "P-Value") %>% <math>str_extract(., "(?<=,[:space:]).*")
# r_rand <- str_subset(test, "Costes Shuffled Mean") %>%
   str_extract(., "(?<=,[:space:]).*")
#
# sd_rand <- str_subset(test, "Costes Shuffled Std") %>%
   str_extract(., "(?<=,[:space:]).*")
\# res <- c(all\_of(sample), all\_of(m1), all\_of(m2), all\_of(r\_pearson),
            all_of(Pval), all_of(r_rand), all_of(sd_rand))
# pgp_coloc2 <- rbind(pgp_coloc2, res)</pre>
#pgp_coloc2 <- pgp_coloc2 %>% na.omit(pgp_coloc2) %>%
# mutate(sex = as.factor(str_sub(sample, start=1L, end=1L)),
         mouseID = as.factor(str_extract(sample, "(?<=-)[:digit:]+")),</pre>
        #prep = as.factor(str_extract(sample, "[:lower:]{2,}")),
#
        prep = as.factor(str extract(sample, "wm/section")),
         image = str_extract(sample, "[:alpha:]{2,}.*"), .before=m1)
#pgp_coloc2$prep
```

```
#pgp_coloc2 <- pgp_coloc2 %>% select(-sample) %>%
# mutate(m1 = as.double(m1),
#
        m2 = as.double(m2),
#
        r pearson = as.double(r pearson),
        Pval = as.double(Pval),
#
#
        r rand = as.double(r rand),
#
        sd_rand = as.double(sd_rand),
        coloc = Pval > 0.95)
#write.xlsx(pgp_coloc2, "../data/IHC/coloc2-pgp-totals.xlsx")
#starting from the processed excel sheet
pgp_coloc2 <- readxl::read_excel("../data/IHC/coloc2-pgp-totals.xlsx") %>%
 mutate(coloc = as.logical(coloc),
        sex = as.factor(sex),
        mouseID = as.factor(mouseID),
        prep = as.factor(prep))
head(pgp_coloc2)
## # A tibble: 6 x 11
          mouseID prep image
                                            m2 r_pea~1 Pval r_rand sd_rand coloc
   sex
    <fct> <fct> <fct> <fct> <fct> <
                                    <dbl> <dbl>
                                                 <dbl> <dbl> <dbl>
                                                                      <dbl> <lgl>
                                                                          O TRUE
## 1 F
          87
                        wm-20x-1 0.01 0.047
                                                  0.21
                                                               0.01
                  wm
                                                           1
## 2 F
          87
                        wm-20x-2 0.021 0.044
                                                  0.28
                                                           1 -0.01
                                                                          O TRUE
                  wm
## 3 F
         87
                        wm-20x-3 0.655 0.585
                                                  0.33
                                                                          0 TRUE
                  wm
                                                              0
                                                           1
## 4 F
         87
                       wm-20x-4
                                   0.719 0.479
                                                  0.38
                                                          1 -0.01
                                                                          0 TRUE
                  wm
        87
                        wm-20x-5
                                                                          0 TRUE
## 5 F
                                   0.674 0.331
                                                  0.3
                                                          1 0
                  wm
         10
## 6 M
                  wm
                        GFP-wm-20x~ 0.012 0.04
                                                  0.13
                                                          1 -0.01
                                                                         O TRUE
## # ... with abbreviated variable name 1: r_pearson
```

### Bcrp

```
#starting with the raw data
#bcrp_filelist <- list.files(path="../data/IHC/coloc2_results_bcrp/",
#
                             pattern=".*.txt", full.names = TRUE)
#bcrp_cols <- c("sample", "m1", "m2", "r_pearson", "Pval", "r_rand", "sd_rand")
#bcrp_coloc2 <- as.data.frame(matrix(,1,length(bcrp_cols)))</pre>
#names(bcrp_coloc2) <- bcrp_cols</pre>
#for (i in bcrp_filelist) {
# test <- read_lines(i)</pre>
# sample <- str_subset(test, "Working on") %>%
   str_extract(., "(?<=:[:space:]).*")
# r_pearson <- str_subset(test, "no threshold") %>%
  str_extract(., "(?<=,[:space:]).*")
# m1 <- str_subset(test, "tM1") %>% str_extract(., "(?<=,[:space:]).*")
# m2 <- str_subset(test, "tM2") %>% str_extract(., "(?<=,[:space:]).*")</pre>
\# Pval \leftarrow str_subset(test, "P-Value") %>% str_extract(., "(?<=,[:space:]).*")
# r rand <- str subset(test, "Costes Shuffled Mean") %>%
# str_extract(., "(?<=,[:space:]).*")
```

```
# sd_rand <- str_subset(test, "Costes Shuffled Std") %>%
   str_extract(., "(?<=,[:space:]).*")
# print(sample)
\# res <- c(all_of(sample), all_of(m1), all_of(m2), all_of(r_pearson),
            all_of(Pval), all_of(r_rand), all_of(sd_rand))
# bcrp_coloc2 <- rbind(bcrp_coloc2, res)</pre>
#}
#bcrp_coloc2 <- bcrp_coloc2 %>% na.omit(bcrp_coloc2) %>%
# mutate(sex = as.factor(str_sub(sample, start=1L, end=1L)),
         mouseID = as.factor(str_extract(sample, "(?<=-)[:digit:]+")),</pre>
         prep = as.factor(str_extract(sample, "[:lower:]{2,}")),
#
         image = str_extract(sample, "[:alpha:]{2,}.*"), .before=m1)
#
#bcrp_coloc2 <- bcrp_coloc2 %>% select(-sample) %>%
# mutate(m1 = as.double(m1),
#
        m2 = as.double(m2),
#
         r_pearson = as.double(r_pearson),
#
        Pval = as.double(Pval),
        r_rand = as.double(r_rand),
#
#
         sd_rand = as.double(sd_rand),
        coloc = Pval > 0.95)
#write.xlsx(bcrp_coloc2, "../data/IHC/coloc2-bcrp-totals.xlsx")
#starting from processed excel file
bcrp_coloc2 <- readxl::read_excel("../data/IHC/coloc2-bcrp-totals.xlsx") %>%
  mutate(coloc = as.logical(coloc),
         sex = as.factor(sex),
         mouseID = as.factor(mouseID),
         prep = as.factor(prep))
head(bcrp_coloc2)
## # A tibble: 6 x 11
          mouseID prep
                                              m2 r_pea~1 Pval r_rand sd_rand coloc
     sex
                           image
                                        m1
##
     <fct> <fct> <fct>
                           <chr>
                                     <dbl> <dbl>
                                                   <dbl> <dbl> <dbl>
                                                                        <dbl> <lgl>
## 1 F
           22
                  section section1~ 0.618 0.229
                                                    0.32
                                                                   0
                                                                            0 TRUE
                                                             1
## 2 F
           22
                 section section1~ 0.787 0.359
                                                    0.63
                                                                    0
                                                                             O TRUE
                 section section1~ 0.782 0.29
## 3 F
           22
                                                    0.51
                                                                    0
                                                                            O TRUE
                                                             1
## 4 F
           22
                 section section3~ 0.517 0.461
                                                    0.49
                                                             1
                                                                    0
                                                                             O TRUE
```

# Mrp2

## 5 F

## 6 F

22

22

0.58

0.58

1

1

section section3~ 0.687 0.541

section section3~ 0.555 0.431

## # ... with abbreviated variable name 1: r\_pearson

O TRUE

0 TRUE

0

0

```
#mrp2_coloc2 <- as.data.frame(matrix(,1,length(mrp2_cols)))</pre>
#names(mrp2_coloc2) <- mrp2_cols</pre>
#for (i in mrp2_filelist) {
# test <- read_lines(i)</pre>
# sample <- str_subset(test, "Working on") %>%
   str_extract(., "(?<=:[:space:]).*")
# r pearson <- str subset(test, "no threshold") %>%
    str extract(., "(?<=,[:space:]).*")
#
# m2 <- str_subset(test, "tM2") %>% str_extract(., "(?<=,[:space:]).*")
# Pval <- str_subset(test, "P-Value") %>% str_extract(., "(?<=,[:space:]).*")
# r rand <- str subset(test, "Costes Shuffled Mean") %>%
   str_extract(., "(?<=,[:space:]).*")
# sd_rand <- str_subset(test, "Costes Shuffled Std") %>%
  str_extract(., "(?<=,[:space:]).*")
# res \leftarrow c(all_of(sample), all_of(m1), all_of(m2), all_of(r_pearson),
#
           all_of(Pval), all_of(r_rand), all_of(sd_rand))
# mrp2_coloc2 <- rbind(mrp2_coloc2, res)</pre>
#}
#mrp2_coloc2 <- mrp2_coloc2 %>% na.omit(mrp2_coloc2) %>%
# mutate(sex = as.factor(str_sub(sample, start=1L, end=1L)),
          mouseID = as.factor(str_extract(sample, "(?<=-)[:digit:]+")),</pre>
#
          image = str extract(sample, "s.*(?=)"),
#
          roi = str_extract(sample, "(?<=_).*"), .before=m1)</pre>
#mrp2_coloc2 <- mrp2_coloc2 %>% select(-sample) %>%
# mutate(m1 = as.double(m1),
#
       m2 = as.double(m2),
#
        r_pearson = as.double(r_pearson),
#
        Pval = as.double(Pval),
        r_rand = as.double(r_rand),
#
#
        sd_rand = as.double(sd_rand),
        coloc = Pval > 0.95)
#write.xlsx(mrp2_coloc2, "../data/IHC/coloc2-mrp2-totals.xlsx")
#starting with processed excel file
mrp2_coloc2 <- readxl::read_excel("../data/IHC/coloc2-mrp2-totals.xlsx") %>%
 mutate(coloc = as.logical(coloc),
        sex = as.factor(sex),
        mouseID = as.factor(mouseID))
head(mrp2_coloc2)
## # A tibble: 6 x 11
    sex mouseID image
                                            m2 r_pea~1 Pval r_rand sd_rand coloc
                              roi
                                      m1
    <fct> <fct> <chr>
                              <chr> <dbl> <dbl> <dbl> <dbl> <dbl> <
                                                                      <dbl> <lgl>
## 1 F
          22
                section2-4~ gcl1 0.063 0.36
                                                  0.12 1
                                                              -0.01
                                                                      0.04 TRUE
                section2-4~ inl1 0
                                                              0
## 2 F
          22
                                                 -0.01 0.34
                                                                      0.02 FALSE
                                         0.087
                                                                      0.02 FALSE
## 3 F
          22
                section2-4~ inl2 0
                                         0.097 -0.03 0.12
                                                             0
```

```
section2-4~ gcl1 0.111 0.001
                                                                  0.01 FALSE
## 4 F
          22
                                             0.01 0.77 0
## 5 F
          22
                 section2-4~ inl1 0 0.183
                                              -0.02 0.18 0
                                                                  0.03 FALSE
## 6 F
          22
                 section2-4~ gcl1 0
                                       0.181
                                              -0.02 0.14 0
                                                                  0.02 FALSE
## # ... with abbreviated variable name 1: r_pearson
```

# Generating summary statistics

```
P-gp
pgp_sumstats <- pgp_coloc2 %>% summarise(m1_mean = mean(m1),
                     m1 sd = sd(m1),
                     m2_{mean} = mean(m2),
                     m2 sd = sd(m2),
                     r_mean = mean(r_pearson),
                     r_{sd} = sd(r_{pearson}),
                     n_{total} = n(),
                     n coloc = sum(coloc)
                     )
pgp_sumstats
## # A tibble: 1 x 8
    m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total n_coloc
##
       <dbl> <dbl> <dbl> <dbl> <dbl> <int> <int>
      0.253 0.294 0.195 0.236 0.280 0.139
                                                   41
bcrp sumstats <- bcrp coloc2 %>% summarise(m1 mean = mean(m1),
                     m1_sd = sd(m1),
                     m2 mean = mean(m2),
                     m2_sd = sd(m2),
                     r_mean = mean(r_pearson),
                     r_{sd} = sd(r_{pearson}),
                     n_{total} = n(),
                     n_{coloc} = sum(coloc),
                     mean_P = mean(Pval)
                     )
bcrp_sumstats
## # A tibble: 1 x 9
    m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total n_coloc mean_P
       <dbl> <dbl> <dbl> <dbl> <dbl> <int>
                                                         <int> <dbl>
## 1 0.737 0.132 0.266 0.0870 0.460 0.104
                                                            42
mrp2_sumstats <- mrp2_coloc2 %>% summarise(m1_mean = mean(m1),
                     m1_sd = sd(m1),
                     m2_{mean} = mean(m2),
                     m2_sd = sd(m2),
                     r_mean = mean(r_pearson),
                     r_{sd} = sd(r_{pearson}),
                     n_{total} = n(),
                     n_{coloc} = sum(coloc)
                     )
mrp2_sumstats
```

```
## # A tibble: 1 x 8
    <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <
                                             <int>
      0.110 0.144
                   0.208 0.157 0.0350 0.0713
                                               145
## 1
                                                       55
mrp2_sumstats_image <- mrp2_coloc2 %>% group_by(image,mouseID,sex,coloc) %>%
 summarise(m1\_mean = mean(m1),
           m1_sd = sd(m1),
           m2_{mean} = mean(m2),
           m2_sd = sd(m2),
           r_mean = mean(r_pearson),
           r_{sd} = sd(r_{pearson}),
           n total = n(),
           n_{coloc} = sum(coloc)
## 'summarise()' has grouped output by 'image', 'mouseID', 'sex'. You can override
## using the '.groups' argument.
mrp2_sumstats_image
## # A tibble: 59 x 12
## # Groups: image, mouseID, sex [40]
##
     image
             mouseID sex coloc m1_mean
                                         m1_sd m2_mean
                                                        m2_sd r_mean
                                                                        r_sd
##
     <chr>
             <fct>
                     <fct> <lgl>
                                 <dbl>
                                         <dbl> <dbl>
                                                        <dbl>
                                                               <dbl>
                                                                        <dbl>
                     F
                          FALSE 0.0527 0.0352 0.0933 0.107
## 1 section~ 40
                                                              0.0167 0.00577
                                               0.204 0.0566 -0.055
## 2 section~ 92
                    М
                          FALSE 0
                                        0
                                                                      0.0778
## 3 section~ 40
                          FALSE 0.155 NA
                                                0.023 NA
                    F
                                                              0.02
                                                       0.0465 0.0533 0.00577
## 4 section~ 40
                   F
                         TRUE
                                 0.127
                                        0.0396 0.129
## 5 section~ 92
                   M FALSE 0.019
                                        0.0329 0.213
                                                       0.0721 0
                                                                      0.0346
## 6 section~ 40
                   F FALSE 0
                                                0.079 NA
                                                              0
                                                                     NA
                                       NA
                    F TRUE
## 7 section~ 40
                                 0.0465 0.0573 0.102
                                                      0.0983 0.025
                                                                     0.00707
## 8 section~ 92
                    M
                          FALSE 0
                                       NA
                                                0.352 NA
                                                              -0.02
                                                                     NΑ
## 9 section~ 92
                          TRUE
                                 0.308
                                        0.0629 0.258
                                                       0.0643 0.11
                                                                     0.0283
## 10 section~ 40
                    F
                          FALSE 0.013 NA
                                                0.362 NA
                                                              0.02
## # ... with 49 more rows, and 2 more variables: n_total <int>, n_coloc <int>
pgp_sumstats_animals <- pgp_coloc2 %>% group_by(mouseID,sex,prep) %>%
 summarise(m1_mean = mean(m1),
           m1_sd = sd(m1),
           m2 mean = mean(m2),
           m2 sd = sd(m2),
           r_mean = mean(r_pearson),
           r_{sd} = sd(r_{pearson}),
           n_{total} = n(),
           n_{coloc} = sum(coloc)
```

## 'summarise()' has grouped output by 'mouseID', 'sex'. You can override using
## the '.groups' argument.

```
pgp_sumstats_animals
```

```
## # A tibble: 8 x 11
## # Groups:
              mouseID, sex [8]
                          m1 mean
                                   m1_sd m2_mean m2_sd r_mean r_sd n_total
    mouseID sex
                  prep
            <fct> <fct>
                            <dbl>
                                    <dbl>
                                          <dbl> <dbl> <dbl> <dbl> <dbl>
     <fct>
## 1 10
                           0.01 0.00784 0.043 0.0168 0.128 0.0295
            М
                  wm
## 2 11
            М
                  wm
                           0.0258 0.0409
                                         0.0503 0.0192 0.145 0.0191
## 3 80
            Μ
                  section 0.320 0.282
                                          0.246 0.290 0.37 0.140
## 4 81
            М
                  section 0.283 0.251
                                          0.204 0.286
                                                         0.377 0.0838
                                                                           6
                  section 0.381 0.325
## 5 82
                                          0.208 0.270
                                                         0.352 0.183
                                                                           6
            М
## 6 83
            М
                           0.430 0.373
                                         0.368 0.274
                                                        0.328 0.0804
                                                                           5
                  wm
## 7 87
                                          0.297 0.247
            F
                  wm
                           0.416 0.366
                                                         0.3
                                                               0.0628
                                                                           5
## 8 9
                           0.0265 0.0155 0.0812 0.0538 0.138 0.0419
            Μ
                  wm
## # ... with 1 more variable: n_coloc <int>
bcrp_sumstats_animals <- bcrp_coloc2 %>% group_by(mouseID,sex,prep) %>%
  summarise(m1_mean = mean(m1),
            m1_sd = sd(m1),
            m2 mean = mean(m2),
            m2 sd = sd(m2),
            r_mean = mean(r_pearson),
            r_{sd} = sd(r_{pearson}),
            n_{total} = n(),
            n_{coloc} = sum(coloc),
           mean_P = mean(Pval)
            )
## 'summarise()' has grouped output by 'mouseID', 'sex'. You can override using
## the '.groups' argument.
bcrp_sumstats_animals
## # A tibble: 7 x 12
              mouseID, sex [7]
## # Groups:
                          m1 mean m1 sd m2 mean m2 sd r mean r sd n total
##
    mouseID sex
                  prep
     <fct> <fct> <fct>
                           <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <
                  section 0.658 0.114
                                                        0.518 0.110
## 1 22
                                          0.385 0.115
            F
                                                                          6
                  section 0.774 0.0717
## 2 40
            F
                                         0.274 0.0454 0.502 0.117
                                                                          6
## 3 80
            Μ
                 section 0.775 0.0587 0.162 0.0183 0.443 0.0638
                                                                          6
## 4 81
                  section 0.663 0.127
                                          0.276 0.0642 0.42 0.103
            М
                                                                          6
                  section 0.754 0.0375 0.275 0.0144 0.408 0.0337
## 5 82
                                                                          6
            Μ
                  section 0.806 0.0483 0.264 0.0486 0.503 0.0403
## 6 92
            М
                                                                          6
## 7 98
            F
                  section 0.732 0.280
                                          0.226 0.0874 0.428 0.177
                                                                          6
## # ... with 2 more variables: n_coloc <int>, mean_P <dbl>
mrp2_sumstats_animals <- mrp2_coloc2 %>% group_by(mouseID,sex,coloc) %>%
  summarise(m1_mean = mean(m1),
            m1 sd = sd(m1),
            m2_{mean} = mean(m2),
            m2_sd = sd(m2),
```

```
r_mean = mean(r_pearson),
            r_{sd} = sd(r_{pearson}),
            n_{total} = n(),
            n_{coloc} = sum(coloc)
            )
## 'summarise()' has grouped output by 'mouseID', 'sex'. You can override using
## the '.groups' argument.
uneven <- mrp2_sumstats_animals %>% group_by(mouseID) %>%
  summarise(n = n(), coloc = coloc, sex=sex) %>%
  filter(n==1) %>% select(-n) %>%
  mutate(coloc = toString(as.integer(coloc))) %>%
  mutate(coloc = str_replace(toString(coloc), pattern = "0", replacement = "1")) %>%
  mutate(coloc = as.logical(as.integer(coloc)))
## 'summarise()' has grouped output by 'mouseID'. You can override using the
## '.groups' argument.
mrp2_sumstats_animals <- rbind(mrp2_sumstats_animals, uneven) %>% arrange(., mouseID)
mrp2_sumstats_animals
## # A tibble: 16 x 11
              mouseID, sex [8]
## # Groups:
     mouseID sex
                   coloc m1 mean
                                     m1_sd m2_mean
                                                     m2\_sd
                                                                       r_sd
                                                             r_{mean}
##
                            <dbl>
                                                     <dbl>
                                                              <dbl>
      <fct>
             <fct> <lgl>
                                     <dbl> <dbl>
                                                                       <dbl>
## 1 22
             F
                   FALSE 0.0234
                                   0.0497
                                            0.202
                                                   0.118 -0.0262
                                                                     0.0421
## 2 22
             F
                   TRUE
                          0.04
                                   0.0325
                                                   0.164
                                                            0.08
                                            0.244
                                                                     0.0566
## 3 40
             F
                   FALSE 0.0543
                                   0.0658
                                            0.115
                                                    0.117
                                                            0.0122
                                                                     0.0109
## 4 40
             F
                   TRUE
                          0.0796
                                   0.0552
                                            0.136
                                                    0.0669 0.0486
                                                                     0.0177
## 5 62
                   FALSE 0.0655
                                   0.0959
                                            0.0723 0.0747 -0.0174
                                                                     0.0254
             М
## 6 62
                   TRUE NA
                                  NA
                                           NA
                                                   NA
                                                           NA
                                                                    NA
## 7 77
             M
                   FALSE 0.035
                                   0.0492
                                            0.135
                                                    0.0385 0.00933
                                                                     0.0249
## 8 77
             М
                   TRUE
                          0.242
                                   0.142
                                            0.200
                                                    0.0504 0.08
                                                                      0.0486
## 9 81
                   FALSE 0.00646 0.00785 0.434
             М
                                                    0.239 -0.00385 0.0112
## 10 81
                   TRUE
                          0.029
                                   0.0428
                                            0.420
                                                    0.291
                                                            0.044
                                                                     0.0313
             М
## 11 82
             М
                   FALSE 0.0468
                                   0.0604
                                            0.143
                                                    0.0994 0.024
                                                                     0.0397
## 12 82
             Μ
                   TRUE
                          0.326
                                   0.170
                                            0.218
                                                    0.116
                                                            0.128
                                                                     0.0669
## 13 92
             М
                   FALSE 0.0406
                                   0.0896
                                            0.198
                                                    0.103 -0.0167
                                                                     0.0424
## 14 92
             М
                   TRUE
                          0.306
                                   0.0802
                                            0.306
                                                    0.0831 0.115
                                                                     0.03
## 15 98
             F
                   FALSE 0.056
                                   0.0752
                                            0.251
                                                    0.0982 0.005
                                                                     0.0404
             F
## 16 98
                   TRUE
                          0.253
                                   0.124
                                            0.262
                                                    0.0856 0.131
                                                                     0.0752
## # ... with 2 more variables: n_total <int>, n_coloc <int>
pgp_sumstats_sex <- pgp_coloc2 %>% group_by(sex) %>%
  summarise(m1\_mean = mean(m1),
            m1_sd = sd(m1),
            m2_{mean} = mean(m2),
            m2_sd = sd(m2),
            r mean = mean(r pearson),
```

 $r_{sd} = sd(r_{pearson}),$ 

```
n_{total} = n(),
           n_{coloc} = sum(coloc)
           )
pgp_sumstats_sex
## # A tibble: 2 x 9
   sex m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total n_coloc
    <fct> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <int> <int>
## 1 F
           0.416 0.366 0.297 0.247 0.3 0.0628
                                                  5
                                                          5
## 2 M
           36
                                                          36
bcrp_sumstats_sex <- bcrp_coloc2 %>% group_by(sex) %>%
 summarise(m1_mean = mean(m1),
           m1 sd = sd(m1),
           m2 mean = mean(m2),
           m2_sd = sd(m2),
           r_mean = mean(r_pearson),
           r_sd = sd(r_pearson),
           n_{total} = n(),
           n_{coloc} = sum(coloc)
bcrp_sumstats_sex
## # A tibble: 2 x 9
## sex m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total n_coloc
   <fct> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <int> <int>
## 1 F
                        0.295 0.107 0.483 0.136
                                                            18
          0.721 0.176
                                                    18
## 2 M
          mrp2_sumstats_sex <- mrp2_coloc2 %>% group_by(sex) %>%
 summarise(m1_mean = mean(m1),
           m1 sd = sd(m1),
           m2_{mean} = mean(m2),
           m2\_sd = sd(m2),
           r_mean = mean(r_pearson),
           r_{sd} = sd(r_{pearson}),
           n_{total} = n(),
           n coloc = sum(coloc)
           )
mrp2_sumstats_sex
## # A tibble: 2 x 9
   sex m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total n_coloc
## <fct> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <int> <int>
           ## 1 F
                                                  53
                                                          24
## 2 M
          0.113 0.154 0.212 0.178 0.0314 0.0664
                                                   92
                                                          31
pgp_sumstats_prep <- pgp_coloc2 %>% group_by(prep) %>%
 summarise(m1\_mean = mean(m1),
           m1_sd = sd(m1),
           m2 mean = mean(m2),
```

```
m2_sd = sd(m2),
            r_mean = mean(r_pearson),
            r_{sd} = sd(r_{pearson}),
            n_{total} = n(),
            mean_P = mean(Pval)
pgp_sumstats_prep
## # A tibble: 2 x 9
    prep m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total mean_P
           <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <int> <dbl>
    <fct>
## 1 section 0.328 0.273 0.219 0.266 0.366 0.133
                                                      18
             ## 2 wm
mrp2_sumstats_coloc <- mrp2_coloc2 %>% group_by(coloc) %>%
 summarise(m1\_mean = mean(m1),
           m1_sd = sd(m1),
           m2_{mean} = mean(m2),
           m2_sd = sd(m2),
           r_mean = mean(r_pearson),
           r_{sd} = sd(r_{pearson}),
           n_{total} = n(),
           mean_P = mean(Pval)
mrp2_sumstats_coloc
## # A tibble: 2 x 9
    coloc m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total mean_P
    <lgl> <dbl> <dbl> <dbl> <dbl> <
                                      <dbl> <dbl> <int> <dbl>
## 1 FALSE 0.0393 0.0672 0.187 0.166 -0.00622 0.0329
                                                       90 0.431
## 2 TRUE 0.225 0.160 0.243 0.136 0.102 0.0656
                                                        55 0.996
```

# Statistical Tests

## difference in location

0.2141667

##

```
coin::wilcox_test(pgp_sumstats_animals$r_mean ~ pgp_sumstats_animals$prep, conf.int=TRUE, distribution=

##

## Exact Wilcoxon-Mann-Whitney Test

##

## data: pgp_sumstats_animals$r_mean by pgp_sumstats_animals$prep (section, wm)

## Z = 2.2361, p-value = 0.03571

## alternative hypothesis: true mu is not equal to 0

## 95 percent confidence interval:

## 0.02366667 0.24866667

## sample estimates:
```

```
coin::wilcox_test(pgp_sumstats_animals$m1_mean ~ pgp_sumstats_animals$prep, conf.int=TRUE, distribution
##
## Exact Wilcoxon-Mann-Whitney Test
## data: pgp_sumstats_animals$m1_mean by pgp_sumstats_animals$prep (section, wm)
## Z = 0.44721, p-value = 0.7857
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.1473 0.3710
## sample estimates:
## difference in location
##
                 0.25675
coin::wilcox_test(pgp_sumstats_animals$m2_mean ~ pgp_sumstats_animals$prep, conf.int=TRUE, distribution
##
## Exact Wilcoxon-Mann-Whitney Test
##
## data: pgp_sumstats_animals$m2_mean by pgp_sumstats_animals$prep (section, wm)
## Z = 0.44721, p-value = 0.7857
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.1636000 0.2033333
## sample estimates:
## difference in location
##
               0.1264167
coin::wilcox_test(pgp_sumstats_animals$r_mean ~ pgp_sumstats_animals$sex, conf.int=TRUE, distribution="
## Warning in cci(alpha): cannot compute confidence interval
## Exact Wilcoxon-Mann-Whitney Test
## data: pgp_sumstats_animals$r_mean by pgp_sumstats_animals$sex (F, M)
## Z = -0.21822, p-value = 1
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
##
                  -0.028
coin::wilcox_test(pgp_sumstats_animals$m1_mean ~ pgp_sumstats_animals$sex, conf.int=TRUE, distribution=
## Warning in cci(alpha): cannot compute confidence interval
## Exact Wilcoxon-Mann-Whitney Test
```

```
##
## data: pgp_sumstats_animals$m1_mean by pgp_sumstats_animals$sex (F, M)
## Z = 1.0911, p-value = 0.5
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
##
                   0.1333
coin::wilcox_test(pgp_sumstats_animals$m2_mean ~ pgp_sumstats_animals$sex, conf.int=TRUE, distribution=
## Warning in cci(alpha): cannot compute confidence interval
##
  Exact Wilcoxon-Mann-Whitney Test
##
## data: pgp_sumstats_animals$m2_mean by pgp_sumstats_animals$sex (F, M)
## Z = 1.0911, p-value = 0.5
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
                   0.0932
##
pgp_wm_sumstats <- pgp_sumstats_animals %>% filter(prep == "wm")
coin::wilcox_test(pgp_wm_sumstats$r_mean ~ pgp_wm_sumstats$sex, conf.int=TRUE, distribution="exact")
## Warning in cci(alpha): cannot compute confidence interval
##
## Exact Wilcoxon-Mann-Whitney Test
## data: pgp_wm_sumstats$r_mean by pgp_wm_sumstats$sex (F, M)
## Z = 0.70711, p-value = 0.8
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
                  0.15875
coin::wilcox_test(pgp_wm_sumstats$m1_mean ~ pgp_wm_sumstats$sex, conf.int=TRUE, distribution="exact")
## Warning in cci(alpha): cannot compute confidence interval
## Exact Wilcoxon-Mann-Whitney Test
##
```

```
## data: pgp_wm_sumstats$m1_mean by pgp_wm_sumstats$sex (F, M)
## Z = 0.70711, p-value = 0.8
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
                                                        0.389675
coin::wilcox_test(pgp_wm_sumstats$m2_mean ~ pgp_wm_sumstats$sex, conf.int=TRUE, distribution="exact")
## Warning in cci(alpha): cannot compute confidence interval
##
         Exact Wilcoxon-Mann-Whitney Test
##
## data: pgp_wm_sumstats$m2_mean by pgp_wm_sumstats$sex (F, M)
## Z = 0.70711, p-value = 0.8
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
                                                           0.23145
pgp_sect_sumstats <- pgp_sumstats_animals %>% filter(prep == "section")
#coin::wilcox_test(pqp_sect_sumstats$r_mean ~ pqp_sect_sumstats$sex, conf.int=TRUE, distribution="exact
\#coin::wilcox\_test(pgp\_sect\_sumstats\$m1\_mean \sim pgp\_sect\_sumstats\$sex, conf.int=TRUE, distribution="exact properties of the properties of
\#coin::wilcox\_test(pgp\_sect\_sumstats\$m2\_mean \sim pgp\_sect\_sumstats\$sex, conf.int=TRUE, distribution="exact properties of the properties of
coin::wilcox_test(bcrp_sumstats_animals$r_mean ~ bcrp_sumstats_animals$sex, conf.int=TRUE, distribution
## Warning in cci(alpha): cannot compute confidence interval
##
## Exact Wilcoxon-Mann-Whitney Test
##
## data: bcrp_sumstats_animals$r_mean by bcrp_sumstats_animals$sex (F, M)
## Z = 1.0607, p-value = 0.4
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
                                                 0.03916667
coin::wilcox_test(bcrp_sumstats_animals$m1_mean ~ bcrp_sumstats_animals$sex, conf.int=TRUE, distribution
## Warning in cci(alpha): cannot compute confidence interval
```

```
##
## Exact Wilcoxon-Mann-Whitney Test
##
## data: bcrp_sumstats_animals$m1_mean by bcrp_sumstats_animals$sex (F, M)
## Z = -1.0607, p-value = 0.4
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
##
                   -0.027
coin::wilcox_test(bcrp_sumstats_animals$m2_mean ~ bcrp_sumstats_animals$sex, conf.int=TRUE, distribution
## Warning in cci(alpha): cannot compute confidence interval
##
## Exact Wilcoxon-Mann-Whitney Test
## data: bcrp_sumstats_animals$m2_mean by bcrp_sumstats_animals$sex (F, M)
## Z = 0.35355, p-value = 0.8571
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
               0.03708333
mrp2_sumstats_animals$coloc <- as.factor(mrp2_sumstats_animals$coloc)</pre>
mrp2_sumstats_animals$r_mean <- replace_na(mrp2_sumstats_animals$r_mean, 0)
mrp2_sumstats_animals$m1_mean <- replace_na(mrp2_sumstats_animals$m1_mean, 0)
mrp2_sumstats_animals$m2_mean <- replace_na(mrp2_sumstats_animals$m2_mean, 0)</pre>
coin::wilcoxsign_test(r_mean ~ coloc | mouseID, data=mrp2_sumstats_animals, conf.int=TRUE, distribution
## Exact Wilcoxon-Pratt Signed-Rank Test
##
## data: y by x (pos, neg)
## stratified by block
## Z = -2.5205, p-value = 0.007812
## alternative hypothesis: true mu is not equal to 0
coin::wilcoxsign_test(m1_mean ~ coloc | mouseID, data=mrp2_sumstats_animals, conf.int=TRUE, distribution
##
## Exact Wilcoxon-Pratt Signed-Rank Test
## data: y by x (pos, neg)
   stratified by block
## Z = -1.9604, p-value = 0.05469
## alternative hypothesis: true mu is not equal to 0
```

```
coin::wilcoxsign_test(m2_mean ~ coloc | mouseID, data=mrp2_sumstats_animals, conf.int=TRUE, distribution
##
##
  Exact Wilcoxon-Pratt Signed-Rank Test
## data: y by x (pos, neg)
## stratified by block
## Z = -1.4003, p-value = 0.1953
## alternative hypothesis: true mu is not equal to 0
mrp2_pos_sumstats_animals <- mrp2_sumstats_animals %>% filter(coloc== TRUE)
coin::wilcox_test(mrp2_pos_sumstats_animals$r_mean ~ mrp2_pos_sumstats_animals$sex, conf.int=TRUE, dist
##
## Exact Wilcoxon-Mann-Whitney Test
##
\verb|## data: mrp2_pos_sumstats_animals$r_mean by mrp2_pos_sumstats_animals$sex (F, M)
## Z = 0.59987, p-value = 0.6071
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.06642857 0.08733333
## sample estimates:
## difference in location
              0.004571429
coin::wilcox_test(mrp2_pos_sumstats_animals$m1_mean ~ mrp2_pos_sumstats_animals$sex, conf.int=TRUE, dis
##
## Exact Wilcoxon-Mann-Whitney Test
## data: mrp2_pos_sumstats_animals$m1_mean by mrp2_pos_sumstats_animals$sex (F, M)
## Z = -0.14907, p-value = 1
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.2864667 0.2530667
## sample estimates:
## difference in location
              -0.05243333
coin::wilcox_test(mrp2_pos_sumstats_animals$m2_mean ~ mrp2_pos_sumstats_animals$sex, conf.int=TRUE, dis
##
## Exact Wilcoxon-Mann-Whitney Test
## data: mrp2_pos_sumstats_animals$m2_mean by mrp2_pos_sumstats_animals$sex (F, M)
## Z = -0.14907, p-value = 1
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.2846286 0.2623333
## sample estimates:
## difference in location
             -0.04391667
##
```

# Plotting the data

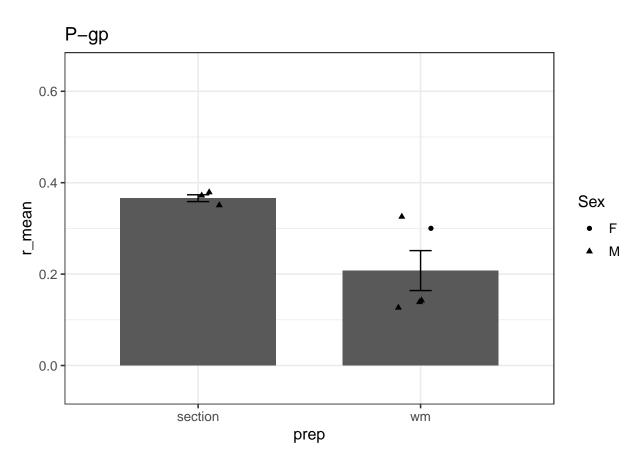
theme\_bw(12) +

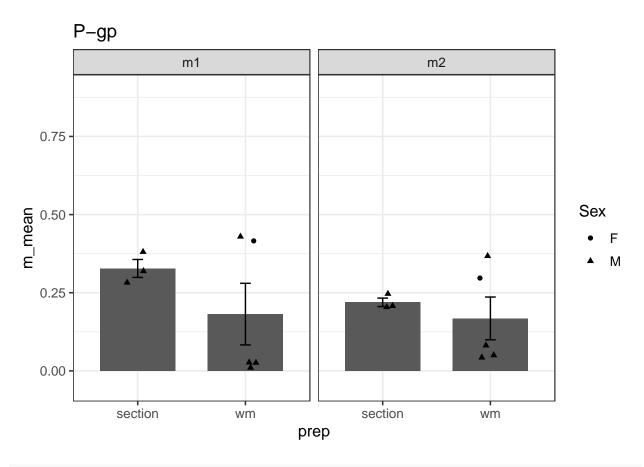
pgp\_animal\_r\_plot

ylim(-0.05,0.65) + ggtitle("P-gp")

guides(shape=guide\_legend("Sex")) +

```
pgp_coloc2$coloc <- as.logical(pgp_coloc2$coloc)</pre>
pgp_coloc2$mouseID <- as.factor(pgp_coloc2$mouseID)</pre>
pgp_coloc2_long <- pgp_coloc2 %>% pivot_longer(cols = m1:m2, names_to = "m_stat_type", values_to = "m_s
pgp_coloc2_long <- pgp_coloc2_long %>% group_by(mouseID, m_stat_type, prep, sex) %>%
  summarise(m_mean = mean(m_stat_val),
             m_sd = sd(m_stat_val),
             r_mean = mean(r_pearson),
             r_{sd} = sd(r_{pearson}),
             n_{total} = n(),
             n_{coloc} = sum(coloc))
## 'summarise()' has grouped output by 'mouseID', 'm_stat_type', 'prep'. You can
## override using the '.groups' argument.
#per animal
pgp_animal_r_plot <- pgp_sumstats_animals %>%
  ggplot(aes(x=prep, y=r_mean)) +
  stat_summary(fun = 'mean', geom="bar", width=0.7) +
  stat_summary(fun.data = mean_se,
                 geom = "errorbar", width=0.1) +
  geom_jitter(color="black", width=0.1, aes(shape=sex)) +
```

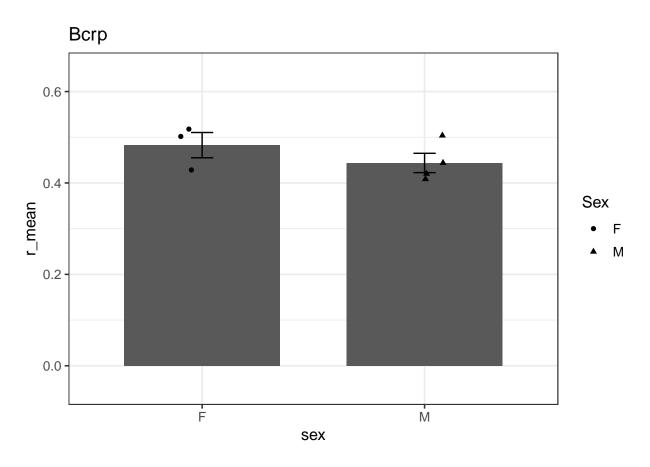




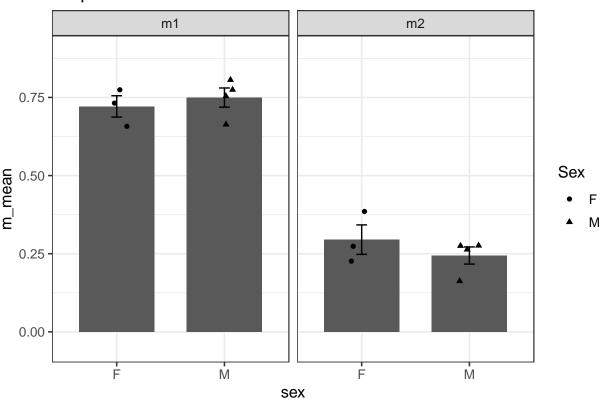
 $\#ggsave("../figures/pgp\_m\_plot.png", plot=pgp\_animal\_m\_plot, width=5, height=3)$ 

## 'summarise()' has grouped output by 'mouseID', 'm\_stat\_type'. You can override
## using the '.groups' argument.

```
ggtitle("Bcrp")
bcrp_animal_r_plot
```







#ggsave("../figures/bcrp\_m\_plot.png", plot=bcrp\_animal\_m\_plot, width=5, height=3)

## 'summarise()' has grouped output by 'mouseID', 'coloc', 'm\_stat\_type', 'sex'.
## You can override using the '.groups' argument.

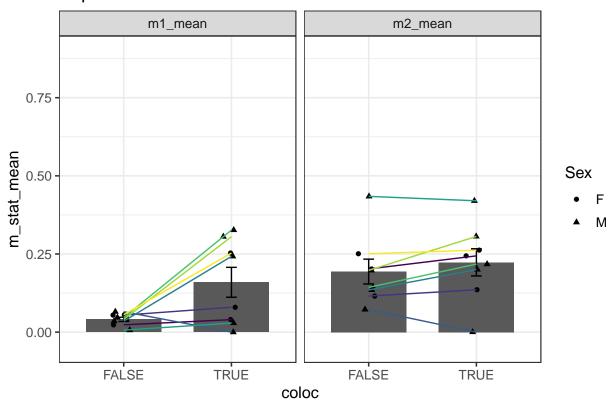
```
mrp2_sumstats_animals_long <- mrp2_sumstats_animals %>% pivot_longer(cols = c(m1_mean,m2_mean), names_t
#per animal
mrp2_animal_r_plot <- mrp2_sumstats_animals %>%
    ggplot(aes(x=coloc, y=r_mean)) +
    stat_summary(fun = 'mean', geom="bar", width=0.7) +
    stat_summary(fun.data = mean_se,
```

```
geom = "errorbar", width=0.1) +
geom_jitter(color="black", width=0.1, aes(shape=sex)) +
geom_line(aes(group=mouseID, color=mouseID)) +
scale_color_viridis(discrete="TRUE") +
theme_bw(12) +
guides(shape=guide_legend("Sex"), color = "none") +
ylim(-0.1,0.65) +
ggtitle("Mrp2")
mrp2_animal_r_plot
```

# Mrp2 0.4 0.4 0.2 0.0 FALSE COloc TRUE

```
ggtitle("Mrp2")
mrp2_animal_m_plot
```

# Mrp2



##ggsave("../figures/mrp2\_m\_plot.png", plot=mrp2\_animal\_m\_plot, width=5, height=3)