Transporter (P-gp) Colocalization Analysis (ImageJ Coloc2)

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

Reading in the data

P-gp

```
pgp_filelist <- list.files(path="../data/IHC/coloc2_results_pgp/", pattern="*.txt", full.names = TRUE)
pgp_cols <- c("sample", "m1", "m2", "r_pearson", "Pval", "r_rand", "sd_rand")</pre>
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:]).*")</pre>
  r_pearson <- str_subset(test, "no threshold") %>% str_extract(., "(?<=,[:space:]).*")
  m1 <- str_subset(test, "tM1") %>% str_extract(., "(?<=,[:space:]).*")</pre>
  m2 <- str_subset(test, "tM2") %>% str_extract(., "(?<=,[:space:]).*")</pre>
  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:]).*")</pre>
 res <- c(all_of(sample), all_of(m1), all_of(m2), all_of(r_pearson), all_of(Pval), all_of(r_rand), all
 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,}")),
        image = str_extract(sample, "[:alpha:]{2,}.*"), .before=m1)
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") head(pgp_coloc2)
```

```
sex mouseID prep
                                            m2 r pearson Pval r rand sd rand coloc
                             image
## 2
      F
                                                                 0.01
                                                                            O TRUE
              87
                          wm-20x-1 0.010 0.047
                                                     0.21
                   wm
                                                             1
## 3
       F
              87
                          wm-20x-2 0.021 0.044
                                                     0.28
                                                                -0.01
                                                                            0
                                                                               TRUE
                   wm
## 4
                                                                 0.00
                                                                            O TRUE
       F
              87
                   wm
                          wm-20x-3 0.655 0.585
                                                     0.33
                                                             1
## 5
              87
                          wm-20x-4 0.719 0.479
                                                     0.38
                                                             1 -0.01
                                                                            0 TRUE
                   wm
## 6
                          wm-20x-5 0.674 0.331
                                                                            0 TRUE
       F
              87
                                                     0.30
                                                                 0.00
                   wm
                                                             1
## 7
              10
                  wm GFP-wm-20x-1 0.012 0.040
                                                     0.13
                                                             1 -0.01
                                                                            O TRUE
```

Bcrp

```
bcrp_filelist <- list.files(path="../data/IHC/coloc2_results_bcrp/", pattern=".*.txt", full.names = TRU
bcrp_cols <- c("sample", "m1", "m2", "r_pearson", "Pval", "r_rand", "sd_rand")</pre>
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:]).*")</pre>
  r_pearson <- str_subset(test, "no threshold") %>% str_extract(., "(?<=,[:space:]).*")
  m1 <- str_subset(test, "tM1") %>% str_extract(., "(?<=,[:space:]).*")</pre>
  m2 <- str_subset(test, "tM2") %>% str_extract(., "(?<=,[:space:]).*")</pre>
  Pval <- str_subset(test, "P-Value") %% str_extract(., "(?<=,[:space:]).*")</pre>
  r_rand <- str_subset(test, "Costes Shuffled Mean") %>% str_extract(., "(?<=,[:space:]).*")
  sd_rand <- str_subset(test, "Costes Shuffled Std") %>% str_extract(., "(?<=,[:space:]).*")</pre>
  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
  bcrp_coloc2 <- rbind(bcrp_coloc2, res)</pre>
}
```

```
## [1] "F-22-section1-40x-1-avg4"
## [1] "F-22-section1-40x-2-avg4"
## [1] "F-22-section1-40x-3-avg4"
## [1] "F-22-section3-40x-1-avg4"
## [1] "F-22-section3-40x-2-avg4"
## [1] "F-22-section3-40x-3-avg4"
## [1] "F-GTL-section1-40x-1-avg4"
## [1] "F-GTL-section1-40x-2-avg4"
## [1] "F-GTL-section1-40x-3-avg4"
## [1] "F-GTL-section3-40x-1-avg4"
## [1] "F-GTL-section3-40x-2-avg4"
## [1] "F-GTL-section3-40x-3-avg4"
## [1] "F-McKO-98-section1-40x-1-avg4"
## [1] "F-McKO-98-section1-40x-2-avg4"
## [1] "F-McKO-98-section1-40x-3-avg4"
## [1] "F-McKO-98-section3-40x-1-avg4"
## [1] "F-McKO-98-section3-40x-2-avg4"
## [1] "F-McKO-98-section3-40x-3-avg4"
```

```
## [1] "M-80-section1-40x-1-avg4"
## [1] "M-80-section1-40x-2-avg4"
## [1] "M-80-section1-40x-3-avg4"
## [1] "M-80-section3-40x-1-avg4"
## [1] "M-80-section3-40x-2-avg4"
## [1] "M-80-section3-40x-3-avg4"
## [1] "M-81-section1-40x-1-avg4"
## [1] "M-81-section1-40x-2-avg4"
## [1] "M-81-section1-40x-3-avg4"
## [1] "M-81-section3-40x-1-avg4"
## [1] "M-81-section3-40x-2-avg4"
## [1] "M-81-section3-40x-3-avg4"
## [1] "M-82-section1-40x-1-avg4"
## [1] "M-82-section1-40x-2-avg4"
## [1] "M-82-section1-40x-3-avg4"
## [1] "M-82-section3-40x-1-avg4"
## [1] "M-82-section3-40x-2-avg4"
## [1] "M-82-section3-40x-3-avg4"
## [1] "M-McKO-92-section1-40x-1-avg4"
## [1] "M-McKO-92-section1-40x-2-avg4"
## [1] "M-McKO-92-section1-40x-3-avg4"
## [1] "M-McKO-92-section3-40x-1-avg4"
## [1] "M-McKO-92-section3-40x-2-avg4"
## [1] "M-McKO-92-section3-40x-3-avg4"
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")
head(bcrp_coloc2)
##
     sex mouseID
                                        image
                                                       m2 r_pearson Pval r_rand
                                                 m1
## 2
              22 section section1-40x-1-avg4 0.618 0.229
                                                               0.32
                                                                        1
                                                                               0
## 3
              22 section section1-40x-2-avg4 0.787 0.359
                                                               0.63
                                                                               0
              22 section section1-40x-3-avg4 0.782 0.290
## 4
       F
                                                               0.51
                                                                               0
                                                                        1
## 5
              22 section section3-40x-1-avg4 0.517 0.461
                                                               0.49
              22 section section3-40x-2-avg4 0.687 0.541
## 6
                                                               0.58
                                                                        1
                                                                               0
## 7
       F
              22 section section3-40x-3-avg4 0.555 0.431
                                                               0.58
                                                                               0
##
     sd_rand coloc
           O TRUE
           O TRUE
## 3
```

```
## 4 O TRUE
## 5 O TRUE
## 6 O TRUE
## 7 O TRUE
```

Mrp2

```
mrp2_filelist <- list.files(path="../data/IHC/coloc2_results_mrp2/", pattern="*.txt", full.names = TRUE</pre>
mrp2_cols <- c("sample", "m1", "m2", "r_pearson", "Pval", "r_rand", "sd_rand")</pre>
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:]).*")</pre>
  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 <- 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:]).*")</pre>
  res <- c(all_of(sample), all_of(m1), all_of(m2), all_of(r_pearson), all_of(Pval), all_of(r_rand), all
  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")
head(mrp2_coloc2)
##
     sex mouseID
                                                  m2 r_pearson Pval r_rand sd_rand
                            image roi
                                            m1
```

```
## 2
             22 section2-40x-1 gcl1 0.063 0.360
                                                     0.12 1.00 -0.01
                                                                         0.04
## 3
     F
             22 section2-40x-1 inl1 0.000 0.087
                                                    -0.01 0.34
                                                                0.00
                                                                         0.02
## 4
             22 section2-40x-1 inl2 0.000 0.097
                                                    -0.03 0.12
                                                                 0.00
                                                                         0.02
     F
## 5
      F
             22 section2-40x-2 gcl1 0.111 0.001
                                                     0.01 0.77
                                                                 0.00
                                                                         0.01
             22 section2-40x-2 inl1 0.000 0.183
                                                                 0.00
## 6
      F
                                                    -0.02 0.18
                                                                         0.03
## 7
             22 section2-40x-3 gcl1 0.000 0.181
                                                    -0.02 0.14
                                                                 0.00
                                                                         0.02
##
    coloc
```

```
## 2 TRUE
## 3 FALSE
## 4 FALSE
## 5 FALSE
## 6 FALSE
## 7 FALSE
```

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
                  m1_sd m2_mean
       m1_mean
                                      m2\_sd
                                                r_{mean}
                                                            r_sd n_total n_coloc
## 1 0.2532927 0.294287 0.195439 0.2355011 0.2804878 0.1387795
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)
                      )
bcrp_sumstats
##
       m1_{mean}
                   m1_sd m2_mean
                                         m2\_sd
                                                              r_sd n_total n_coloc
                                                   r_{mean}
## 1 0.7374762 0.1322315 0.2660952 0.08701775 0.4604762 0.104437
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
```

```
m1 sd m2 mean
                                     m2\_sd
      m1 mean
                                              r_{mean}
                                                           r_sd n_total n_coloc
## 1 0.1098828 0.1436731 0.2081172 0.1569569 0.03496552 0.07127923
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
##
              <fct>
                      <fct> <lgl>
                                  <dbl>
                                          <dbl>
                                                  <dbl>
                                                          <dbl>
                                                                  <dbl>
     <chr>
                                                                           <dbl>
                     F
                           FALSE 0.0527 0.0352 0.0933 0.107
                                                                 0.0167 0.00577
## 1 section~ 40
## 2 section~ 92
                     M
                           FALSE 0
                                         0
                                                 0.204
                                                         0.0566 -0.055
                                                                        0.0778
                    F
                           FALSE 0.155 NA
## 3 section~ 40
                                                 0.023 NA
                                                                 0.02
                                                                       NA
                    F TRUE
## 4 section~ 40
                                  0.127
                                         0.0396 0.129
                                                        0.0465 0.0533 0.00577
## 5 section~ 92
                    M FALSE 0.019
                                        0.0329 0.213 0.0721 0
                                                                        0.0346
                         FALSE 0
                    F
## 6 section~ 40
                                         NA
                                                 0.079 NA
                                                                 0
                    F TRUE
## 7 section~ 40
                                  0.0465 0.0573 0.102
                                                        0.0983 0.025
                                                                       0.00707
## 8 section~ 92
                    M FALSE O
                                                 0.352 NA
                                                                -0.02
                                                                       NA
                                         NA
## 9 section~ 92
                           TRUE
                                        0.0629 0.258
                     M
                                  0.308
                                                        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
              mouseID, sex [8]
## # Groups:
                  prep
    mouseID sex
                          m1 mean
                                   m1_sd m2_mean m2_sd r_mean r_sd n_total
                                           <dbl> <dbl> <dbl> <dbl> <
##
    <fct>
            <fct> <fct>
                            <dbl>
                                    <dbl>
## 1 10
                  wm
                           0.01 0.00784 0.043 0.0168 0.128 0.0295
## 2 11
            М
                           0.0258 0.0409
                                         0.0503 0.0192 0.145 0.0191
                                                                            4
                  wm
                  section 0.320 0.282
## 3 80
                                          0.246 0.290
                                                         0.37 0.140
            М
## 4 81
                  section 0.283 0.251
                                          0.204 0.286
                                                         0.377 0.0838
                                                                            6
            Μ
## 5 82
            М
                  section 0.381 0.325
                                          0.208 0.270
                                                         0.352 0.183
## 6 83
                                          0.368 0.274
                                                         0.328 0.0804
                                                                            5
            М
                  wm
                           0.430 0.373
## 7 87
            F
                           0.416 0.366
                                          0.297 0.247
                                                         0.3
                                                              0.0628
                                                                            5
                  wm
                                                                            4
## 8 9
            М
                           0.0265 0.0155
                                          0.0812 0.0538 0.138 0.0419
                  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)
## 'summarise()' has grouped output by 'mouseID', 'sex'. You can override using the
## '.groups' argument.
bcrp_sumstats_animals
## # A tibble: 7 x 11
## # Groups: mouseID, sex [7]
    mouseID sex
                  prep
                          m1_mean m1_sd m2_mean m2_sd r_mean r_sd n_total
    <fct> <fct> <fct>
                            <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <
## 1 22
            F
                  section
                            0.658 0.114
                                          0.385 0.115
                                                        0.518 0.110
                                                                           6
## 2 40
            F
                  section 0.774 0.0717
                                         0.274 0.0454 0.502 0.117
                                                                           6
## 3 80
                  section 0.775 0.0587
                                          0.162 0.0183 0.443 0.0638
            M
                                                                           6
## 4 81
                                          0.276 0.0642 0.42 0.103
            M
                  section
                            0.663 0.127
                                                                           6
## 5 82
                  section 0.754 0.0375
                                          0.275 0.0144 0.408 0.0337
                                                                           6
            М
## 6 92
            Μ
                  section 0.806 0.0483
                                          0.264 0.0486 0.503 0.0403
                                                                           6
## 7 98
            F
                            0.732 0.280
                                          0.226 0.0874 0.428 0.177
                  section
                                                                           6
## # ... with 1 more variable: n_coloc <int>
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.
```

```
mrp2_sumstats_animals
```

```
## # A tibble: 15 x 11
## # Groups:
              mouseID, sex [8]
##
                   coloc m1_mean
                                   m1_sd m2_mean m2_sd
     mouseID sex
                                                          r_{mean}
                                                                   r_sd n_total
##
      <fct>
             <fct> <lgl>
                                   <dbl>
                                           <dbl>
                                                 <dbl>
                                                           <dbl>
                           <dbl>
                                                                 <dbl>
##
   1 22
             F
                   FALSE 0.0234 0.0497
                                          0.202 0.118 -0.0262 0.0421
                                                                             16
##
   2 22
             F
                   TRUE 0.04
                                 0.0325
                                          0.244 0.164
                                                         0.08
                                                                              2
                                                                 0.0566
## 3 40
             F
                   FALSE 0.0543 0.0658
                                          0.115 0.117
                                                         0.0122 0.0109
                                                                              9
## 4 40
             F
                   TRUE 0.0796 0.0552
                                          0.136  0.0669  0.0486  0.0177
                                                                              7
## 5 62
                                          0.0723 0.0747 -0.0174 0.0254
                                                                             19
             М
                   FALSE 0.0655 0.0959
                                          0.135 0.0385 0.00933 0.0249
## 6 77
             М
                   FALSE 0.035
                                 0.0492
                                                                             15
## 7 77
                   TRUE 0.242
                                 0.142
                                          0.200 0.0504
                                                        0.08
                                                                 0.0486
                                                                              7
             М
## 8 81
             М
                   FALSE 0.00646 0.00785 0.434 0.239
                                                        -0.00385 0.0112
                                                                             13
## 9 81
                   TRUE 0.029
                                          0.420 0.291
                                                         0.044
                                                                              5
             М
                                 0.0428
                                                                 0.0313
## 10 82
                   FALSE 0.0468 0.0604
                                          0.143 0.0994 0.024
                                                                 0.0397
                                                                              5
             М
                                          0.218 0.116
                                                                             15
## 11 82
             М
                   TRUE 0.326
                                 0.170
                                                         0.128
                                                                 0.0669
## 12 92
             М
                   FALSE 0.0406 0.0896
                                          0.198 0.103 -0.0167 0.0424
                                                                              9
                                                                              4
## 13 92
             М
                   TRUE 0.306
                                 0.0802
                                          0.306 0.0831 0.115
                                                                 0.03
## 14 98
             F
                   FALSE 0.056
                                 0.0752
                                          0.251 0.0982 0.005
                                                                 0.0404
                                                                              4
             F
## 15 98
                   TRUE 0.253
                                 0.124
                                          0.262 0.0856 0.131
                                                                 0.0752
                                                                             15
## # ... with 1 more variable: n_coloc <int>
pgp_sumstats_sex <- pgp_coloc2 %>% group_by(sex) %>%
 summarise(m1 mean = mean(m1),
            m1 sd = sd(m1),
            m2_{mean} = mean(m2),
```

```
## # A tibble: 2 x 9
           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>
             0.416 0.366
                           0.297 0.247
                                                                   5
## 1 F
                                        0.3
                                              0.0628
                                                           5
## 2 M
             0.231 0.282
                           0.181 0.234 0.278 0.147
                                                          36
                                                                  36
```

```
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> <int> <int>
## 1 F
           18
                                                    18
## 2 M
           24
                                                            24
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> <int> <int>
## 1 F
           0.104 0.124 0.201 0.113 0.0411 0.0794
                                                  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>
## 1 section 0.328 0.273 0.219 0.266 0.366 0.133
                                                    18
                                                           1
             ## 2 wm
                                                    23
                                                           1
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
           <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
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:
## difference in location
##
               0.2141667
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
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
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
```

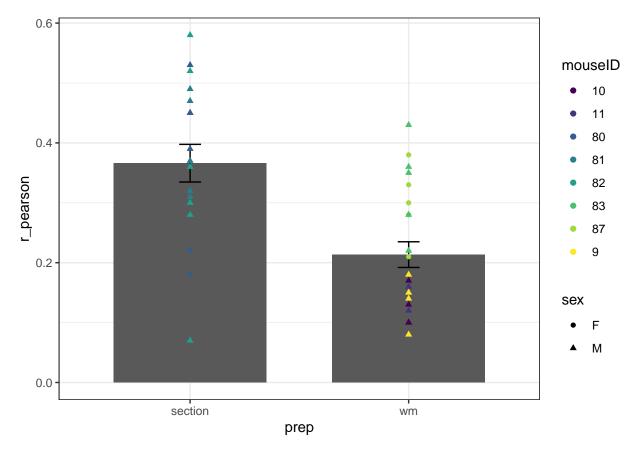
```
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## NA NA
## sample estimates:
## difference in location
##
                 0.23145
coin::wilcox_test(bcrp_sumstats_animals$r_mean ~ bcrp_sumstats_animals$sex, conf.int=TRUE, distribution
## Warning in cci(alpha): 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
```

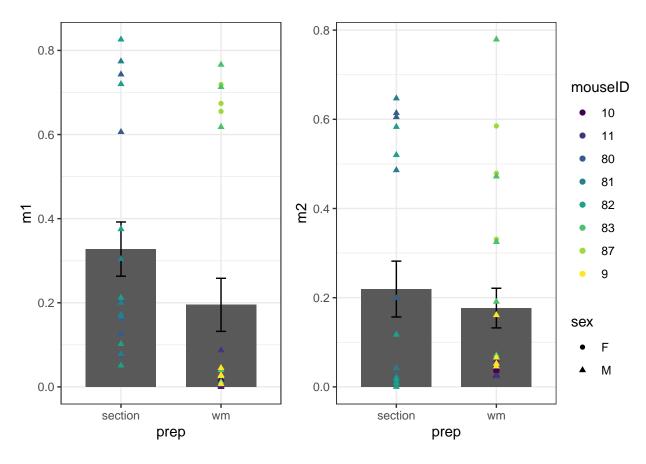
```
## sample estimates:
## difference in location
               0.03708333
mrp2_sumstats_animals$coloc <- as.factor(mrp2_sumstats_animals$coloc)</pre>
coin::wilcox_test(mrp2_sumstats_animals$r_mean ~ mrp2_sumstats_animals$coloc, conf.int=TRUE, distributi
##
## Exact Wilcoxon-Mann-Whitney Test
##
## data: mrp2_sumstats_animals$r_mean by
## mrp2_sumstats_animals$coloc (FALSE, TRUE)
## Z = -3.2433, p-value = 0.0003108
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.1316667 -0.0560000
## sample estimates:
## difference in location
             -0.09666667
coin::wilcox_test(mrp2_sumstats_animals$m1_mean ~ mrp2_sumstats_animals$coloc, conf.int=TRUE, distribut
##
## Exact Wilcoxon-Mann-Whitney Test
## data: mrp2_sumstats_animals$m1_mean by
## mrp2_sumstats_animals$coloc (FALSE, TRUE)
## Z = -1.9674, p-value = 0.05408
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.2649444444 0.0005555556
## sample estimates:
## difference in location
                -0.191719
\#coin::wilcox\_test(mrp2\_sumstats\_animals\$m1\_mean \sim mrp2\_sumstats\_animals\$coloc, conf.int=TRUE)
coin::wilcox_test(mrp2_sumstats_animals$m2_mean ~ mrp2_sumstats_animals$coloc, conf.int=TRUE, distribut
##
## Exact Wilcoxon-Mann-Whitney Test
## data: mrp2_sumstats_animals$m2_mean by
## mrp2_sumstats_animals$coloc (FALSE, TRUE)
## Z = -1.5045, p-value = 0.152
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.17138333 0.03308333
## sample estimates:
## difference in location
##
             -0.06502698
```

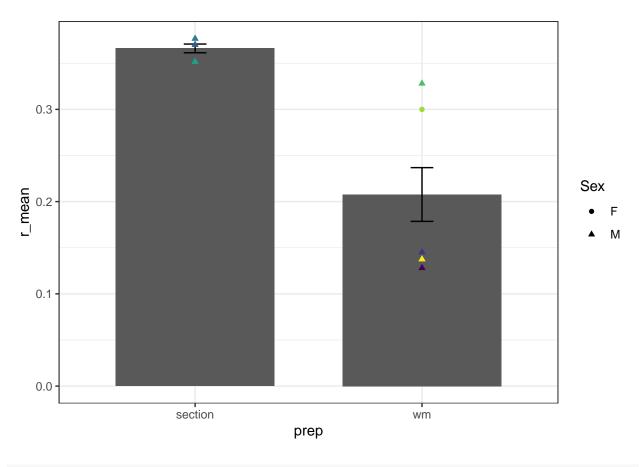
```
##
##
   Exact Wilcoxon-Mann-Whitney Test
## data: mrp2_sumstats_animals$r_mean by mrp2_sumstats_animals$sex (F, M)
## Z = 0.17693, p-value = 0.8881
\#\# alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.06777778 0.06593985
## sample estimates:
## difference in location
##
              0.003111111
coin::wilcox_test(mrp2_sumstats_animals$m1_mean ~ mrp2_sumstats_animals$sex, conf.int=TRUE, distribution
##
##
   Exact Wilcoxon-Mann-Whitney Test
##
## data: mrp2_sumstats_animals$m1_mean by mrp2_sumstats_animals$sex (F, M)
## Z = 0, p-value = 1
## alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.21884821 0.04457143
## sample estimates:
## difference in location
##
              0.00222222
coin::wilcox_test(mrp2_sumstats_animals$m2_mean ~ mrp2_sumstats_animals$sex, conf.int=TRUE, distribution
##
##
   Exact Wilcoxon-Mann-Whitney Test
## data: mrp2_sumstats_animals$m2_mean by mrp2_sumstats_animals$sex (F, M)
## Z = -0.11785, p-value = 0.9546
\#\# alternative hypothesis: true mu is not equal to 0
## 95 percent confidence interval:
## -0.17205128 0.06725833
## sample estimates:
## difference in location
##
              -0.01158512
Plotting the data
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
```

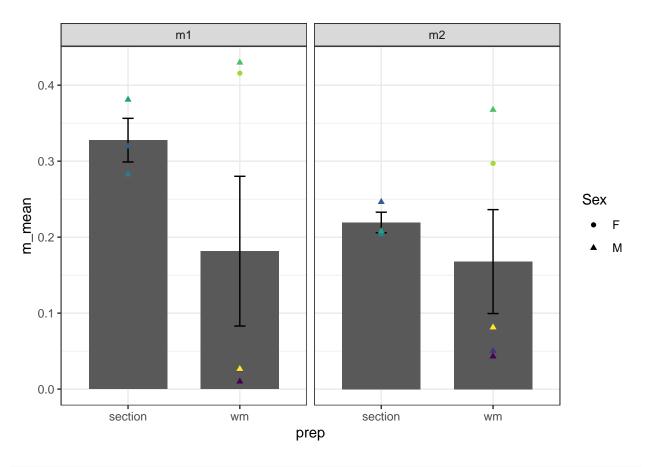
coin::wilcox_test(mrp2_sumstats_animals\$r_mean ~ mrp2_sumstats_animals\$sex, conf.int=TRUE, distribution

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

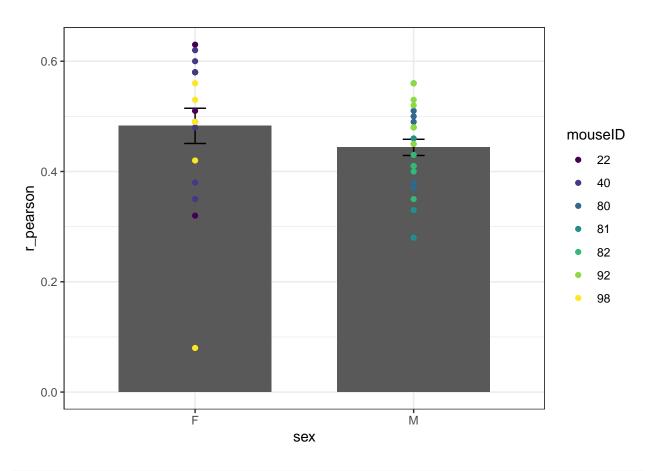


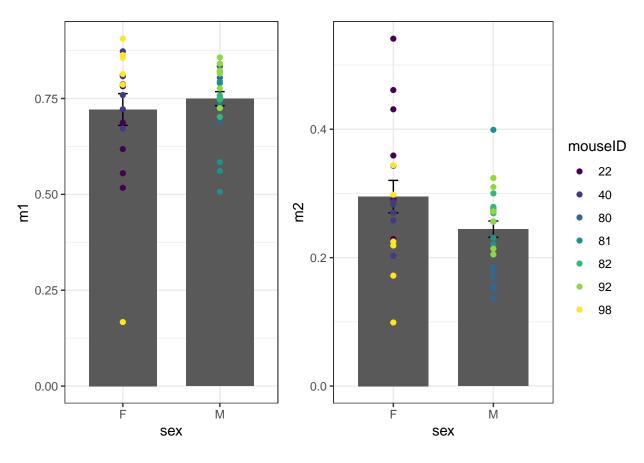


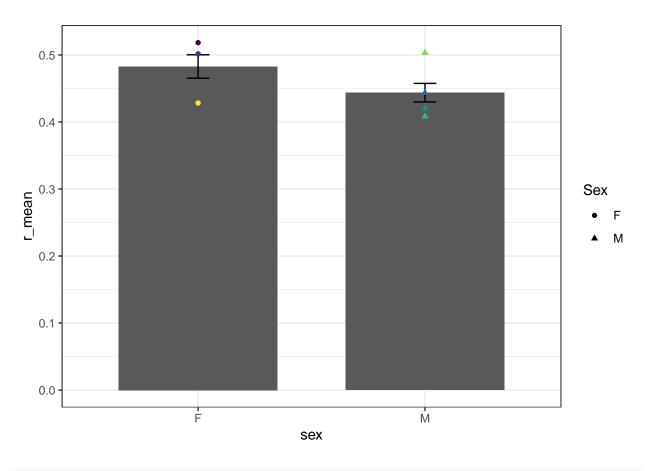


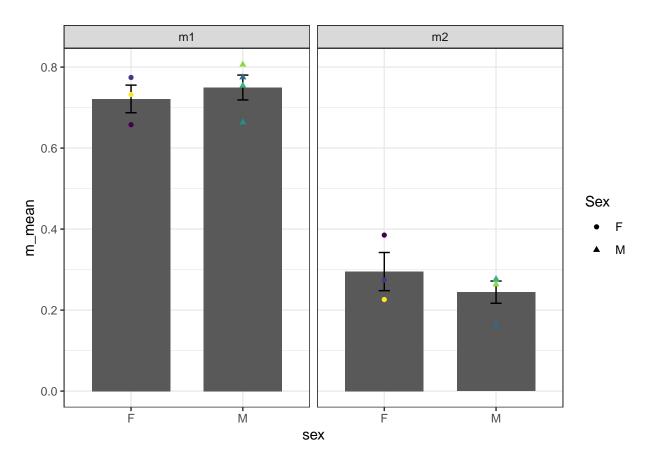


'summarise()' has grouped output by 'mouseID', 'm_stat_type'. You can override
using the '.groups' argument.

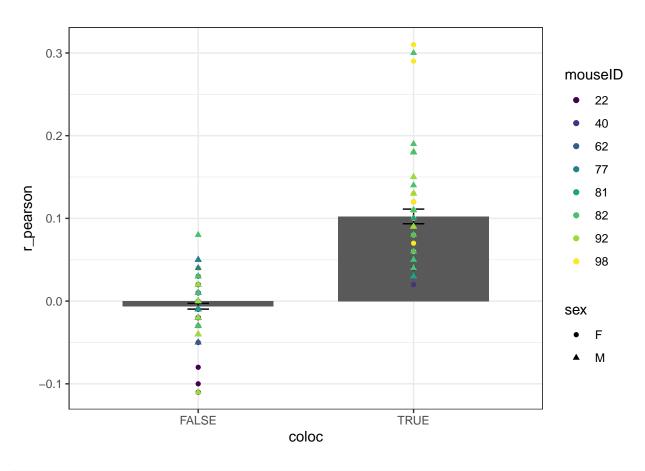


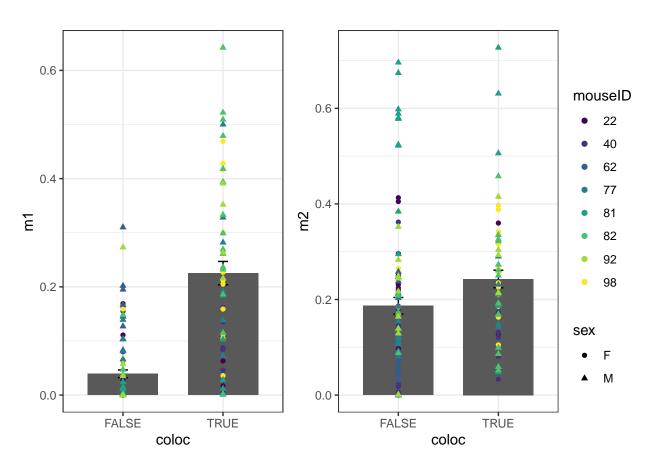


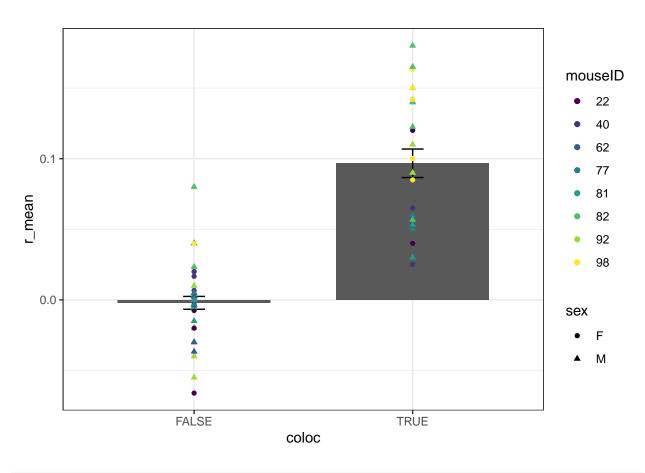


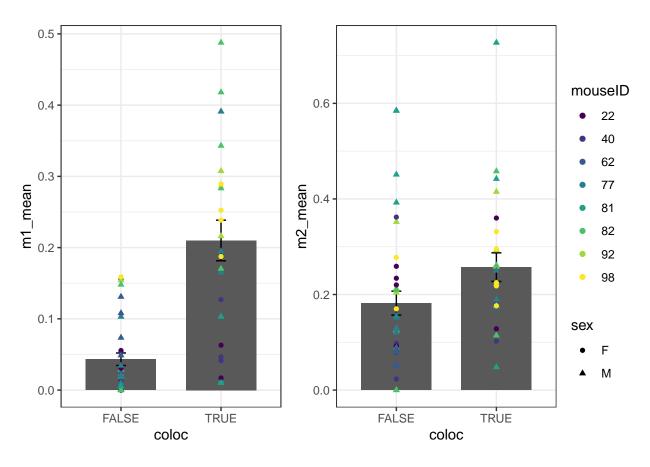


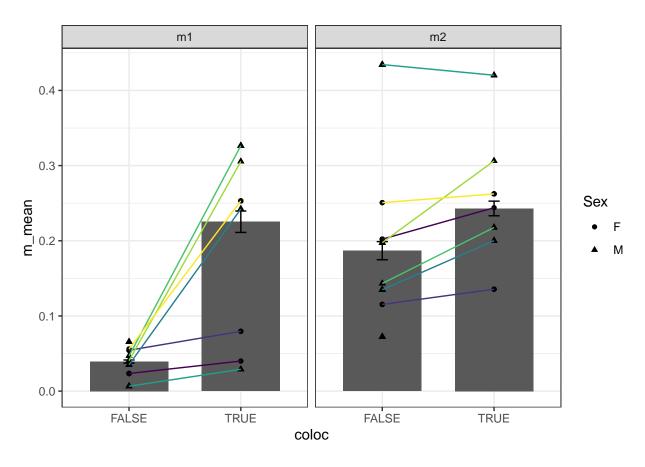
'summarise()' has grouped output by 'mouseID', 'coloc', 'm_stat_type', 'sex'.
You can override using the '.groups' argument.

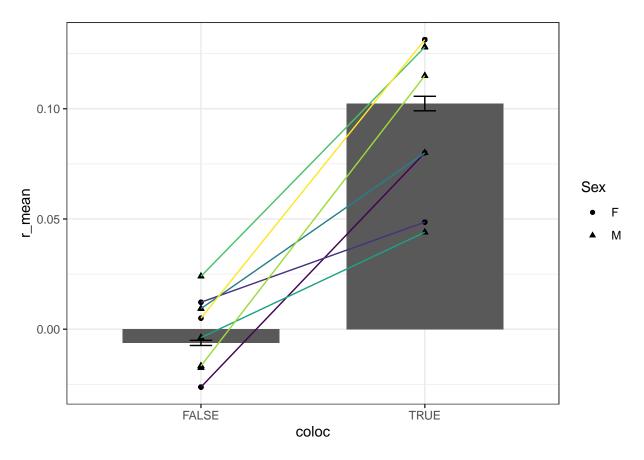












```
pgp_animal_r_plot1 <- pgp_animal_r_plot + ylim(-0.05,0.6) + ggtitle("P-gp")
bcrp_animal_r_plot1 <- bcrp_animal_r_plot + ylim(-0.05,0.6) + ggtitle("Bcrp")</pre>
mrp2_animal_r_plot1 <- mrp2_animal_r_plot + ylim(-0.05,0.6) + ggtitle("Mrp2")</pre>
all_r_plot <- ggarrange(pgp_animal_r_plot1, bcrp_animal_r_plot1, mrp2_animal_r_plot1, ncol=3, common.le
ggsave("../figures/all_r_plot.png", plot=all_r_plot, width=7, height=4)
ggsave("../figures/all_r_plot.svg", plot=all_r_plot, width=7, height=4)
pgp_animal_m1_plot <- ggplot(pgp_sumstats_animals, aes(x=prep, y=m1_mean)) +
  stat_summary(fun = 'mean', geom="bar", width=0.7) +
  stat_summary(fun.data = mean_se,
                 geom = "errorbar", width=0.1) +
  geom_point(aes(color=mouseID, shape=sex)) +
  scale_color_viridis(discrete="TRUE") +
  theme_bw(12) +
  ylim(0,0.9) +
  ggtitle("P-gp") +
  guides(shape=guide_legend("Sex"), color = "none")
bcrp_animal_m1_plot <- ggplot(bcrp_sumstats_animals, aes(x=sex, y=m1_mean)) +
  stat_summary(fun = 'mean', geom="bar", width=0.7) +
  stat_summary(fun.data = mean_se,
                 geom = "errorbar", width=0.1) +
  geom_point(aes(color=mouseID, shape=sex)) +
  scale_color_viridis(discrete="TRUE") +
 theme bw(12) +
```

```
ylim(0,0.9) +
  ggtitle("Bcrp") +
  guides(shape=guide_legend("Sex"), color = "none")
mrp2_animal_m1_plot <- ggplot(mrp2_sumstats_animals, aes(x=coloc, y=m1_mean)) +</pre>
  stat_summary(fun = 'mean', geom="bar", width=0.7) +
  stat_summary(fun.data = mean_se,
                 geom = "errorbar", width=0.1) +
  geom_point(aes(shape=sex)) +
  geom_line(aes(group=mouseID, color=mouseID)) +
  scale_color_viridis(discrete="TRUE") +
  theme_bw(12) +
  ylim(0,0.9) +
  ggtitle("Mrp2") +
  guides(shape=guide_legend("Sex"), color = "none")
all_m1_plot <- ggarrange(pgp_animal_m1_plot, bcrp_animal_m1_plot, mrp2_animal_m1_plot, ncol=3, common.l
ggsave("../figures/all_m1_plot.png", plot=all_m1_plot, width=7, height=4)
ggsave("../figures/all_m1_plot.svg", plot=all_m1_plot, width=7, height=4)
pgp_animal_m2_plot <- ggplot(pgp_sumstats_animals, aes(x=prep, y=m2_mean)) +
  stat_summary(fun = 'mean', geom="bar", width=0.7) +
  stat_summary(fun.data = mean_se,
                 geom = "errorbar", width=0.1) +
  geom_point(aes(color=mouseID, shape=sex)) +
  scale_color_viridis(discrete="TRUE") +
  theme_bw() +
  ylim(0,0.5) +
  ggtitle("P-gp") +
  guides(shape=guide_legend("Sex"), color = "none")
bcrp_animal_m2_plot <- ggplot(bcrp_sumstats_animals, aes(x=sex, y=m2_mean)) +
  stat_summary(fun = 'mean', geom="bar", width=0.7) +
  stat_summary(fun.data = mean_se,
                 geom = "errorbar", width=0.1) +
  geom_point(aes(color=mouseID, shape=sex)) +
  scale_color_viridis(discrete="TRUE") +
  theme_bw() +
  ylim(0,0.5) +
  ggtitle("Bcrp") +
  guides(shape=guide_legend("Sex"), color = "none")
mrp2_animal_m2_plot <- ggplot(mrp2_sumstats_animals, aes(x=coloc, y=m2_mean)) +
  stat_summary(fun = 'mean', geom="bar", width=0.7) +
  stat_summary(fun.data = mean_se,
                 geom = "errorbar", width=0.1) +
  geom_point(aes(shape=sex)) +
  geom_line(aes(group=mouseID, color=mouseID)) +
  scale_color_viridis(discrete="TRUE") +
```

```
theme_bw() +
ylim(0,0.5) +
ggtitle("Mrp2") +
guides(shape=guide_legend("Sex"), color = "none")

all_m2_plot <- ggarrange(pgp_animal_m2_plot, bcrp_animal_m2_plot, mrp2_animal_m2_plot, ncol=3, common.l
ggsave("../figures/all_m2_plot.png", plot=all_m1_plot, width=7, height=4)
ggsave("../figures/all_m2_plot.svg", plot=all_m1_plot, width=7, height=4)</pre>
```