Estimating Viral Prevalence with Data Fusion for Adaptive Two-Phase Pooled Sampling

Initialization

```
# Quicker Simulations
num_replicates <- 20
theta_seq <- seq(0.01, .99, by = .05)

# Settings from Paper, may take a while to compile
#num_replicates <- 200
#theta_seq <- seq(0.01, .99, by = .01)</pre>
```

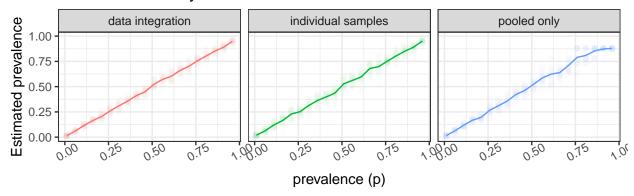
Simulation 1

Part A: N = 100

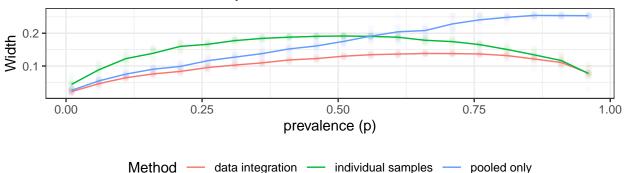
```
set.seed(07012020)
num pooled samples <- 100
num_indiv_samples <- 100</pre>
pool size <- 3
num_theta <- length(theta_seq)</pre>
y_replicates <- array(0, dim = c(num_theta,num_replicates, 2))</pre>
z_replicates <- array(0, dim = c(num_theta,num_replicates, 2))</pre>
yz_replicates <- array(0, dim = c(num_theta,num_replicates, 2))</pre>
y_mean <- y_width <- matrix(0, num_theta, num_replicates)</pre>
z_mean <- z_width <- matrix(0, num_theta, num_replicates)</pre>
yz_mean <- yz_width <- matrix(0, num_theta, num_replicates)</pre>
for (j in 1:num_theta){
  theta <- theta seq[j]
  for (i in 1:num_replicates){
    y <- rbinom(num indiv samples, 1, theta)
    z_mat <- matrix(rbinom(num_pooled_samples * pool_size, 1, theta),</pre>
                     nrow = num_pooled_samples, ncol = pool_size)
    z <- as.numeric(rowSums(z_mat) > 0)
    y_samples <- find_theta_y(y)</pre>
    y_replicates[j,i,] <- y_samples$interval</pre>
    y_mean[j,i] <- y_samples$mean</pre>
    y_width[j,i] <- diff(y_samples$interval)</pre>
    z_samples <- find_theta_z(z_mat, 10000, .1)</pre>
    z_replicates[j,i,] <- z_samples$interval</pre>
```

```
z_mean[j,i] <- mean(z_samples$theta)</pre>
    z_width[j,i] <- diff(z_samples$interval)</pre>
    yz_samples <- find_theta_yz(y,z_mat, 10000, .1)</pre>
    yz_replicates[j,i,] <- yz_samples$interval</pre>
    yz_mean[j,i] <- mean(yz_samples$theta)</pre>
    yz_width[j,i] <- diff(yz_samples$interval)</pre>
}
f1a_dat <- tibble(ci_width = c(y_mean, z_mean, yz_mean),</pre>
         prevalence = rep(theta_seq, 3 * num_replicates),
         Method = rep(c('individual samples', 'pooled only', 'data integration'),
                      each = num_theta * num_replicates))
f1a_mean <- f1a_dat %>%
  group_by(prevalence, Method) %>%
  summarise(width = mean(ci_width),.groups = 'drop')
f1a <- f1a_dat %>%
  ggplot(aes(y=ci_width, x = prevalence, color = Method)) +
  geom_point(alpha = .01) + theme_bw() +
  geom_line(aes(y = width, x = prevalence, color = Method), inherit.aes = F, data = f1a_mean) +
  ggtitle('Posterior Mean by Prevalence') +
  ylab('Estimated prevalence') + xlab("prevalence (p)") +
  facet_wrap(.~Method) + theme(legend.position='none') +
  theme(axis.text.x = element_text(angle = 30))
f1b_dat <- tibble(ci_width = c(y_width, z_width, yz_width),
         prevalence = rep(theta_seq, 3 * num_replicates),
         Method = rep(c('individual samples', 'pooled only', 'data integration'),
                      each = num_theta * num_replicates))
f1b_mean <- f1b_dat %>%
  group_by(prevalence, Method) %>%
  summarise(width = mean(ci_width), .groups = 'drop')
f1b <- f1b_dat %>%
  ggplot(aes(y=ci_width, x = prevalence, color = Method)) +
  geom_point(alpha = .01) + theme_bw() +
  geom_line(aes(y = width, x = prevalence, color = Method), inherit.aes = F, data = f1b_mean) +
  ggtitle('Credible Interval Width by Prevalence') + ylab('Width') +
  xlab("prevalence (p)") + theme(legend.position='bottom')
grid.arrange(f1a,f1b)
```

Posterior Mean by Prevalence



Credible Interval Width by Prevalence

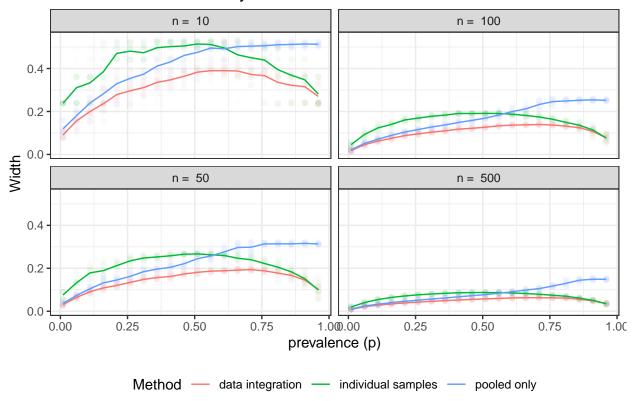


Part B: Varying N

```
num_pooled_samples \leftarrow c(10,50,100,500)
num_indiv_samples <- c(10,50,100,500)
k <- length(num_pooled_samples)</pre>
pool_size <- 3</pre>
num_theta <- length(theta_seq)</pre>
y_replicates <- array(0, dim = c(num_theta,num_replicates, 2,k))</pre>
z_replicates <- array(0, dim = c(num_theta,num_replicates, 2,k))</pre>
yz_replicates <- array(0, dim = c(num_theta,num_replicates, 2,k))</pre>
y_mean <- y_width <- array(0, dim =c(num_theta, num_replicates,k))</pre>
z_mean <- z_width <- array(0, dim = c(num_theta, num_replicates, k))</pre>
yz_mean <- yz_width <- array(0, dim = c(num_theta, num_replicates, k))</pre>
for (j in 1:num_theta){
  theta <- theta_seq[j]</pre>
  for (i in 1:num_replicates){
    for (n_val in 1:k){
      y <- rbinom(num_indiv_samples[n_val], 1, theta)
      z_mat <- matrix(rbinom(num_pooled_samples[n_val] * pool_size, 1, theta),</pre>
                      nrow = num pooled samples[n val], ncol = pool size)
      z <- as.numeric(rowSums(z_mat) > 0)
      y_samples <- find_theta_y(y)</pre>
      y_replicates[j,i,,n_val] <- y_samples$interval</pre>
```

```
y_mean[j,i,n_val] <- y_samples$mean</pre>
      y_width[j,i,n_val] <- diff(y_samples$interval)</pre>
      z_samples <- find_theta_z(z_mat, 10000, .1)</pre>
      z_replicates[j,i,,n_val] <- z_samples$interval</pre>
      z_mean[j,i,n_val] <- mean(z_samples$theta)</pre>
      z_width[j,i,n_val] <- diff(z_samples$interval)</pre>
      yz_samples <- find_theta_yz(y,z_mat, 10000, .1)</pre>
      yz_replicates[j,i,,n_val] <- yz_samples$interval</pre>
      yz_mean[j,i,n_val] <- mean(yz_samples$theta)</pre>
      yz_width[j,i,n_val] <- diff(yz_samples$interval)</pre>
    }
 }
}
f1b_dat <- tibble(ci_width = c(y_width, z_width, yz_width),</pre>
         prevalence = rep(rep(theta_seq, 3 * num_replicates),k),
         n = paste('n = ', rep(rep(num_pooled_samples, each = num_replicates * length(theta_seq)),3)),
         Method = rep(rep(c('individual samples', 'pooled only', 'data integration'),
                       each = num_theta * num_replicates),each =k))
f1b_mean <- f1b_dat %>%
  group_by(prevalence, Method, n) %>%
  summarise(width = mean(ci_width), .groups = 'drop')
f1b_dat %>%
  ggplot(aes(y=ci_width, x = prevalence, color = Method)) +
  geom_point(alpha = .01) + theme_bw() +
  geom_line(aes(y = width, x = prevalence, color = Method), inherit.aes = F, data = f1b_mean) +
  ggtitle('Credible Interval Width by Prevalence') + ylab('Width') +
  xlab("prevalence (p)") + theme(legend.position='bottom') +
  facet_wrap(.~n)
```

Credible Interval Width by Prevalence

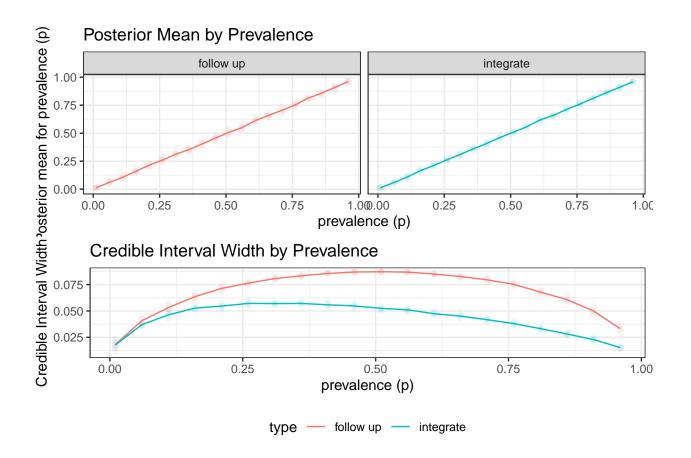


Simulation 2

Part A: fixed n = 100

```
set.seed(08012020)
num_pooled_samples <- 100</pre>
pool_size <- 5</pre>
num_theta <- length(theta_seq)</pre>
## Simulate and Fit Data
followup_mean <- followup_width <- integrate_mean <- integrate_width <-</pre>
  array(0, dim = c(num_theta, num_replicates))
for (j in 1:num_theta){
  theta <- theta_seq[j]</pre>
  for (i in 1:num_replicates){
    y <- rbinom(num_pooled_samples * pool_size, 1, theta)
    z_mat <- matrix(y, nrow = num_pooled_samples, ncol = pool_size)</pre>
    y_tmp <- find_theta_y(y)</pre>
    followup_mean[j,i] <- y_tmp$mean</pre>
    followup_width[j,i] <- diff(y_tmp$interval)</pre>
    followup_samples <- sum(z_mat) * pool_size</pre>
    y2 <- rbinom(followup_samples, 1, theta)</pre>
    yz_tmp <- find_theta_yz(y2, z_mat, num_mcmc = 10000, step_size = .1)</pre>
    integrate_mean[j,i] <- mean(yz_tmp$theta)</pre>
    integrate_width[j,i] <- diff(yz_tmp$interval)</pre>
```

```
}
}
f1_dat <- tibble(ci_mean = c(integrate_mean, followup_mean),</pre>
       replicate = rep(rep(1:num_replicates, each = num_theta),2),
       type = rep(c('integrate', 'follow up'), each = num_theta * num_replicates),
       prevalence = rep(theta_seq, num_replicates * 2))
f1_mean <- f1_dat %>% group_by(type, prevalence) %>%
  summarise(ci_mean= mean(ci_mean), .groups = 'drop')
f1 <- f1_dat %>%
  ggplot(aes(y=ci_mean, x = prevalence, color = type)) +
  geom_point(alpha = .01) +
  geom_line(aes(y = ci_mean, x = prevalence, color = type), inherit.aes = F, data = f1_mean) +
  theme_bw() + ggtitle('Posterior Mean by Prevalence') +
  ylab("Posterior mean for prevalence (p)") + xlab("prevalence (p)") +
  theme(legend.position='none') + facet_wrap(.~type)
f2_dat <- tibble(ci_width = c(integrate_width,followup_width),</pre>
       replicate = rep(rep(1:num_replicates, each = num_theta),2),
       type = rep(c('integrate', 'follow up'), each = num_theta * num_replicates),
       prevalence = rep(theta_seq, num_replicates * 2))
f2_mean <- f2_dat %>% group_by(type, prevalence) %>%
  summarise(ci_width = mean(ci_width), .groups = 'drop')
f2 <- f2_dat %>%
  ggplot(aes(y=ci_width, x = prevalence, color = type)) +
  geom_point(alpha = .01) +
  geom_line(aes(y = ci_width, x = prevalence, color = type), inherit.aes = F, data = f2_mean) +
  theme_bw() + ggtitle('Credible Interval Width by Prevalence') +
  ylab("Credible Interval Width") + xlab("prevalence (p)") +
  theme(legend.position='bottom')
grid.arrange(f1,f2)
```

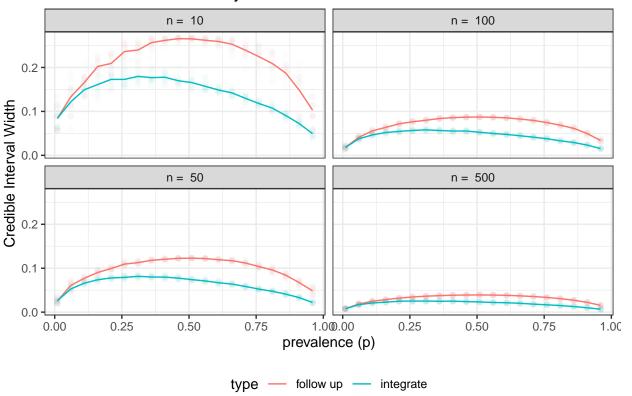


Part B: Varying N

```
set.seed(08012020)
num_pooled_samples <- c(10,50,100,500)
pool_size <- 5</pre>
num_theta <- length(theta_seq)</pre>
k <- length(num_pooled_samples)</pre>
## Simulate and Fit Data
followup_mean <- followup_width <- integrate_mean <- integrate_width <-</pre>
  array(0, dim = c(num theta, num replicates,k))
for (j in 1:num_theta){
  theta <- theta_seq[j]</pre>
  for (i in 1:num replicates){
    for(k2 in 1:k){
      y <- rbinom(num_pooled_samples[k2] * pool_size, 1, theta)
      z_mat <- matrix(y, nrow = num_pooled_samples[k2], ncol = pool_size)</pre>
      y_tmp <- find_theta_y(y)</pre>
      followup_mean[j,i,k2] <- y_tmp$mean</pre>
      followup_width[j,i,k2] <- diff(y_tmp$interval)</pre>
      followup_samples <- sum(z_mat) * pool_size</pre>
      y2 <- rbinom(followup_samples, 1, theta)</pre>
      yz_tmp <- find_theta_yz(y2, z_mat, num_mcmc = 10000, step_size = .1)</pre>
      integrate_mean[j,i,k2] <- mean(yz_tmp$theta)</pre>
      integrate_width[j,i,k2] <- diff(yz_tmp$interval)</pre>
```

```
}
}
f2_dat <- tibble(ci_width = c(integrate_width,followup_width),</pre>
       replicate = rep(rep(rep(1:num_replicates, each = num_theta),2),k),
       type = rep(c('integrate', 'follow up'), each = num_theta * num_replicates * k),
       prevalence = rep(rep(theta_seq, num_replicates * 2),k),
       n = paste('n = ', rep(rep(num_pooled_samples, each = num_theta * num_replicates),2)))
f2_mean <- f2_dat %>% group_by(type, prevalence, n) %>%
  summarise(ci_width = mean(ci_width), .groups = 'drop')
f2_dat %>%
  ggplot(aes(y=ci_width, x = prevalence, color = type)) +
  geom_point(alpha = .01) +
  geom_line(aes(y = ci_width, x = prevalence, color = type), inherit.aes = F, data = f2_mean) +
  theme_bw() + ggtitle('Credible Interval Width by Prevalence') +
  ylab("Credible Interval Width") + xlab("prevalence (p)") +
  theme(legend.position='bottom') + facet_wrap(.~n)
```

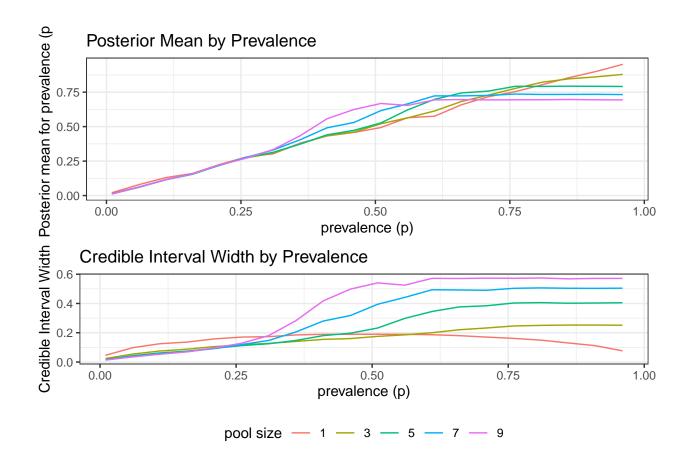
Credible Interval Width by Prevalence



Simulation 3

```
set.seed(07052020)
num_pooled_samples <- 100
pool_size <- c(1,3,5,7,9)
pool_length <- length(pool_size)</pre>
```

```
max_samples <- max(pool_size) * num_pooled_samples</pre>
num_theta <- length(theta_seq)</pre>
## Simulate and Fit Data
z_mean <- z_width <- array(0, dim = c(pool_length,num_theta, num_replicates))</pre>
for (j in 1:num_theta){
  theta <- theta seq[j]
  for (i in 1:num replicates){
    y <- rbinom(max_samples, 1, theta)
    for (k in 1:pool_length){
      z_mat <- matrix(y[1:(pool_size[k]*num_pooled_samples)],</pre>
                      nrow = num_pooled_samples, ncol = pool_size[k])
      z_samples <- find_theta_z(z_mat, 10000, .1)</pre>
      z_mean[k,j,i] <- mean(z_samples$theta)</pre>
      z_width[k,j,i] <- diff(z_samples$interval)</pre>
    }
 }
}
f1_dat <- tibble(ci_mean = c(z_mean),
       replicate = rep(1:num_replicates, each = pool_length * num_theta),
       num_pooled = factor(rep(pool_size, num_replicates * num_theta)),
       prevalence = rep(rep(theta seq, each = pool length), num replicates))
f1_mean <- f1_dat %>% group_by(num_pooled, prevalence) %>%
  summarise(ci_mean = mean(ci_mean), .groups = 'drop')
f1 <- f1_dat %>% ggplot(aes(y=ci_mean, x = prevalence, color = num_pooled)) +
 geom_line(aes(y = ci_mean, x = prevalence, color = num_pooled), inherit.aes = F, data = f1_mean) +
  theme_bw() + ggtitle('Posterior Mean by Prevalence') +
  ylab("Posterior mean for prevalence (p)") +
  xlab("prevalence (p)") + theme(legend.position='none')
f2_dat <- tibble(ci_mean = c(z_width),
       replicate = rep(1:num_replicates, each = pool_length * num_theta),
       `pool size` = factor(rep(pool_size, num_replicates * num_theta)),
       prevalence = rep(rep(theta_seq, each = pool_length),num_replicates))
f2_mean <- f2_dat %>% group_by(`pool size`, prevalence) %>%
  summarise(ci_mean = mean(ci_mean), .groups = 'drop')
f2 <- f2_dat %>% ggplot(aes(y=ci_mean, x = prevalence, color = `pool size`)) +
  geom_line(data = f2_mean) +
  theme_bw() + ggtitle('Credible Interval Width by Prevalence') +
  ylab("Credible Interval Width") + xlab("prevalence (p)") +
  theme(legend.position='bottom')
grid.arrange(f1,f2)
```



Simulation 4

```
### Phase 1. Start with 20 samples of size 5.
# initialize parameters
set.seed(07052020)
theta_length <- length(theta_seq)</pre>
num_theta <- length(theta_seq)</pre>
total_samples <- 100
pool_size <- 5</pre>
z <- array(0, dim = c(num_replicates, theta_length,</pre>
                     total_samples / pool_size, pool_size))
# Take Phase 1 Sample
z_mean <- z_width <- array(0, dim = c(num_replicates, theta_length))</pre>
for (i in 1:num_replicates){
  for (j in 1:theta_length){
   z[i,j,,] <- matrix(rbinom(total_samples,1,theta_seq[j]),</pre>
                      nrow = total_samples / pool_size, ncol = pool_size)
   z_{samples} \leftarrow find_{theta_z(z[i,j,,], 10000, .1)}
    z_mean[i,j] <- mean(z_samples$theta)</pre>
    z_width[i,j] <- diff(z_samples$interval)</pre>
}
```

```
### Phase 2. Take an additional 80 samples.
# a. Add individual samples
phase2_samples <- 80
y width <- y mean <- array(0, dim = c(num replicates, theta length))
for (i in 1:num_replicates){
  for (j in 1:theta_length){
    y <- rbinom(phase2_samples,1,theta_seq[j])</pre>
    y_{\text{samples}} \leftarrow find_{\text{theta}}(y,z[i,j,], 10000, .1)
    y_mean[i,j] <- mean(y_samples$theta)</pre>
    y_width[i,j] <- diff(y_samples$interval)</pre>
  }
}
tib1 <- tibble(vals = c(y_width),
               prevalence = rep(theta_seq, each = num_replicates))
# Add pooled samples of size 3
pool size <- 3
z3_width <- z3_mean <- array(0, dim = c(num_replicates, theta_length))</pre>
for (i in 1:num_replicates){
  for (j in 1:theta_length){
    z3 <- matrix(rbinom(phase2_samples * pool_size,1,theta_seq[j]),</pre>
                 nrow = phase2_samples, ncol = pool_size)
    z_pooled <- as.numeric(c(rowMeans(z[i,j,,]) > 0, rowMeans(z3) > 0))
    n_pooled <- c(rep(5, nrow(z[i,j,,])), rep(pool_size, nrow(z3)))</pre>
    z3_samples <- find_theta_z2(z_pooled, n_pooled, 10000, .1)
    z3_mean[i,j] <- mean(z3_samples$theta)</pre>
    z3_width[i,j] <- diff(z3_samples$interval)</pre>
  }
}
tib3 \leftarrow tibble(vals = c(z3 width),
               prevalence = rep(theta_seq, each = num_replicates))
#Add pooled samples of size 5
pool_size <- 5</pre>
z5_width <- z5_mean <- array(0, dim = c(num_replicates, theta_length))
for (i in 1:num_replicates){
  for (j in 1:theta_length){
    z5 <- matrix(rbinom(phase2_samples * pool_size,1,theta_seq[j]),</pre>
                 nrow = phase2_samples, ncol = pool_size)
    z_pooled <- as.numeric(c(rowMeans(z[i,j,,]) > 0, rowMeans(z5) > 0))
    n_pooled <- c(rep(5, nrow(z[i,j,,])), rep(pool_size, nrow(z5)))</pre>
    z5_samples <- find_theta_z2(z_pooled, n_pooled, 10000, .1)
    z5_mean[i,j] <- mean(z5_samples$theta)</pre>
```

```
z5_width[i,j] <- diff(z5_samples$interval)</pre>
 }
}
tib5 <- tibble(vals = c(z5_width),
               prevalence = rep(theta_seq, each = num_replicates))
f3a_dat \leftarrow tibble(vals = c(c(z_mean), c(y_mean), c(z3_mean), c(z5_mean)),
                   theta = rep(rep(theta_seq, each = num_replicates),4),
                   method = rep(c('phase 1', 'phase 2 with 1', 'phase 2 with 3',
                                   'phase2 with 5'), each = num_replicates * theta_length))
f3a_mean <- f3a_dat %>% group_by(theta, method) %>%
  summarise(vals = mean(vals), .groups = 'drop')
f3a <- f3a_dat %>%
  ggplot(aes(y = vals, x = theta, color = method)) +
  geom_line(aes(y = vals, x = theta, color = method), inherit.aes = F, data = f3a_mean) +
  theme_bw() + ylim(0,1) + ggtitle('Posterior Mean by Strategy') +
  theme(legend.position = "none") + ylab("Posterior mean for prevalence (p)") +
  xlab("prevalence (p)")
f3b_dat \leftarrow tibble(vals = c(c(z_width), c(y_width), c(z3_width), c(z5_width)),
                   theta = rep(rep(theta_seq, each = num_replicates),4),
                   method = rep(c('initial pool','phase 2 with 1', 'phase2 with 3', 'phase2 with 5'),
                                each = num_replicates * theta_length))
f3b_mean <- f3b_dat %>% group_by(theta, method) %>%
  summarise(vals = mean(vals), .groups = 'drop')
f3b <- f3b_dat %>%
  ggplot(aes(y = vals, x = theta, color = method)) + geom_line(data = f3b_mean) +
  theme_bw() + ylim(0,1) + ggtitle('Credible Interval Width by Strategy') +
  theme(legend.position='bottom') + ylab("Credible Interval Width") +
  xlab("prevalence (p)")
grid.arrange(f3a, f3b)
```

