Results simulation study DelayedGSD

April 16, 2024

1 Rejection rate

1.1 2 stages

```
Power by method (columns) and scenario (rows):
                                                                                (nominal level 80%)
                                   fixC ar method 1 method 2 method 3
scenario n.sim missing binding
        1 10000
                    TRUE
                             TRUE FALSE 10
                                              90.43%
                                                        90.37%
                                                                 90.18%
        3 10000
                    TRUE
                             TRUE FALSE 5
                                              90.13%
                                                        90.11%
                                                                 89.78%
        5 10000
                    TRUE
                            TRUE
                                   TRUE 10
                                              90.00%
                                                        90.10%
                                                                 90.18%
        7 10000
                    TRUE
                             TRUE
                                   TRUE
                                              89.73%
                                                        89.79%
                                                                 89.78%
        9
           9900
                    TRUE
                            FALSE
                                   TRUE 10
                                              90.20%
                                                        90.33%
                                                                 90.41%
                                                        90.47%
       11
           9900
                    TRUE
                           FALSE
                                   TRUE
                                              90.31%
                                                                 90.46%
           9900
                                              90.74%
       13
                    TRUE
                           FALSE FALSE 10
                                                        90.67%
                                                                 90.41%
                            FALSE FALSE
           9900
                    TRUE
                                              90.67%
                                                        90.66%
                                                                 90.46%
       17 10000
                             TRUE FALSE
                                              90.38%
                                                                 90.09%
                   FALSE
                                                        90.38%
   Type 1 error by method (columns) and scenario (rows):
                                                                               (nominal level 2.5\%)
scenario n.sim missing binding fixC ar method 1 method 2 method 3
        2 10000
                    TRUE
                             TRUE FALSE 10
                                               2.76%
                                                         2.75%
                                                                  2.68%
        4 10000
                    TRUE
                             TRUE FALSE 5
                                               2.72%
                                                         2.72%
                                                                  2.70%
        6 10000
                    TRUE
                             TRUE
                                   TRUE 10
                                               2.56%
                                                         2.56%
                                                                  2.68%
        8 10000
                    TRUE
                             TRUE
                                   TRUE
                                               2.62%
                                                         2.62%
                                                                  2.70%
       10 10000
                    TRUE
                            FALSE
                                   TRUE 10
                                               2.14%
                                                         2.15%
                                                                  2.33%
       12 10000
                    TRUE
                            FALSE
                                   TRUE
                                               2.29%
                                                         2.30%
                                                                  2.34%
       14 10000
                    TRUE
                                               2.29%
                                                         2.27%
                                                                  2.33%
                           FALSE FALSE 10
       16 10000
                    TRUE
                            FALSE FALSE
                                               2.36%
                                                         2.36%
                                                                  2.34%
       18
           9900
                   FALSE
                             TRUE FALSE
                                               2.70%
                                                         2.70%
                                                                  2.66%
```

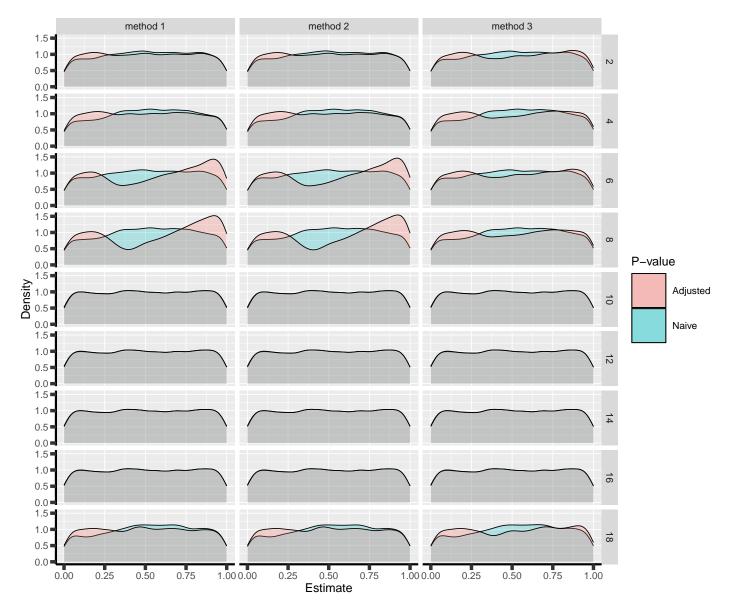


Figure 1: Naive and adjusted p-value distribution over all simulations under the null. Each row correspond to a different scenario

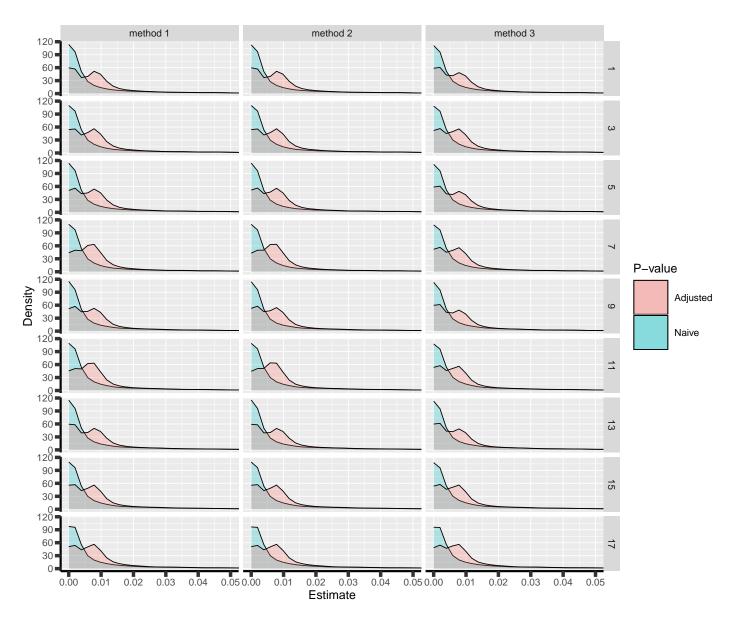


Figure 2: Naive and adjusted p-value distribution over all simulations under the alternative. Each row correspond to a different scenario

1.2 3 stages

18 10000

FALSE

TRUE FALSE 5

Power by method (columns) and scenario (rows): (nominal level 80%) scenario n.sim missing binding fixC ar method 1 method 2 method 3 1 10000 TRUE FALSE 10 TRUE 90.96% 90.85% 90.44% 3 10000 TRUE TRUE FALSE 5 90.62% 90.61% 90.28% 5 10000 TRUE TRUE TRUE 10 90.30% 90.44% 90.44% 7 10000 TRUE TRUE 5 90.21% 90.28% TRUE 90.34% 9 10000 TRUE FALSE TRUE 10 90.01% 90.27% 90.40% 11 10000 FALSE 90.04% TRUE TRUE 89.94% 90.12% 13 10000 TRUE FALSE FALSE 10 90.60% 90.49% 90.40% 15 10000 TRUE FALSE FALSE 5 90.31% 90.31% 90.04% 17 10000 **FALSE** TRUE FALSE 5 89.85% 89.81% 89.57% Type 1 error by method (columns) and scenario (rows): (nominal level 2.5%) scenario n.sim missing binding fixC ar method 1 method 2 method 3 2 10000 TRUE FALSE 10 TRUE 2.46% 2.51% 2.44% 4 10000 2.52% 2.52% 2.49% TRUE TRUE FALSE 5 6 10000 TRUE 10 2.25% 2.24% 2.44% TRUE TRUE 8 10000 TRUE TRUE TRUE 5 2.41% 2.43% 2.49% 10 9931 FALSE TRUE 10 2.30% 2.32% 2.47% TRUE 12 10000 TRUE TRUE FALSE 2.50% 2.51% 2.64% 14 9931 TRUE FALSE FALSE 10 2.45% 2.42% 2.47% 16 10000 FALSE FALSE 5 TRUE 2.52% 2.52% 2.64%

2.65%

2.65%

2.52%

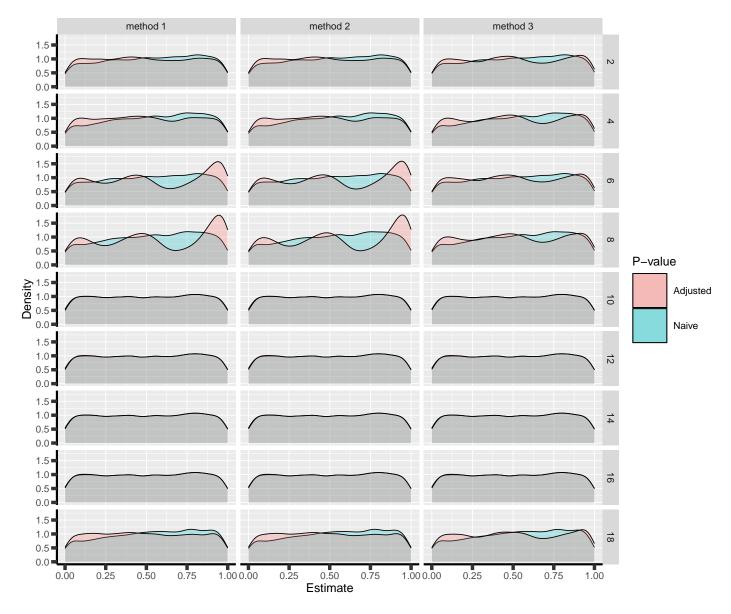


Figure 3: Naive and adjusted p-value distribution over all simulations under the null. Each row correspond to a different scenario

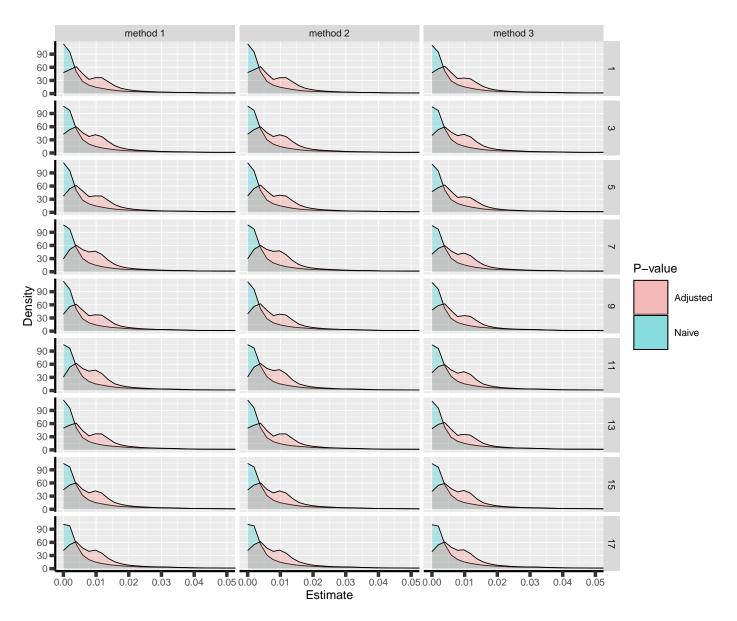


Figure 4: Naive and adjusted p-value distribution over all simulations under the alternative. Each row correspond to a different scenario

2 Conclusion of the trial

2.1 2 stages

Relative frequency of stopping for efficacy/futility at decision/final

• Method 1

	N	missing	hypo	binding	fixC	ar	${\tt decision.eff}$	${\tt decision.fut}$	final.eff	final.fut
1:	10000	TRUE	power	TRUE	FALSE	10	51.23%	2.90%	39.20%	6.67%
2:	10000	TRUE	typeI	TRUE	FALSE	10	0.86%	71.79%	1.90%	25.45%
3:	10000	TRUE	power	TRUE	FALSE	5	48.74%	2.99%	41.39%	6.88%
4:	10000	TRUE	typeI	TRUE	FALSE	5	0.81%	69.49%	1.91%	27.79%
5:	10000	TRUE	power	TRUE	TRUE	10	50.80%	3.33%	39.20%	6.67%
6:	10000	TRUE	typeI	TRUE	TRUE	10	0.66%	71.99%	1.90%	25.45%
7:	10000	TRUE	power	TRUE	TRUE	5	48.34%	3.39%	41.39%	6.88%
8:	10000	TRUE	typeI	TRUE	TRUE	5	0.71%	69.59%	1.91%	27.79%
9:	9900	TRUE	power	FALSE	TRUE	10	50.62%	3.70%	39.59%	6.10%
10:	10000	TRUE	typeI	FALSE	TRUE	10	0.45%	0.23%	1.69%	97.63%
11:	9900	TRUE	power	FALSE	TRUE	5	48.34%	3.47%	41.97%	6.21%
12:	10000	TRUE	typeI	FALSE	TRUE	5	0.55%	0.09%	1.74%	97.62%
13:	9900	TRUE	power	FALSE	FALSE	10	51.15%	3.16%	39.59%	6.10%
14:	10000	TRUE	typeI	FALSE	FALSE	10	0.60%	0.08%	1.69%	97.63%
15:	9900	TRUE	power	FALSE	FALSE	5	48.70%	3.12%	41.97%	6.21%
16:	10000	TRUE	typeI	FALSE	FALSE	5	0.62%	0.02%	1.74%	97.62%
17:	10000	FALSE	power	TRUE	FALSE	5	47.04%	2.46%	43.34%	7.16%
18:	9900	FALSE	typeI	TRUE	FALSE	5	0.82%	67.88%	1.88%	29.42%

Method 2:

	N	missing	hypo	${\tt binding}$	fixC	ar	${\tt decision.eff}$	${\tt decision.fut}$	final.eff	final.fut
1:	10000	TRUE	power	TRUE	FALSE	10	51.29%	3.13%	39.08%	6.50%
2:	10000	TRUE	typeI	TRUE	FALSE	10	0.86%	72.69%	1.89%	24.56%
3:	10000	TRUE	power	TRUE	FALSE	5	48.74%	3.01%	41.37%	6.88%
4:	10000	TRUE	typeI	TRUE	FALSE	5	0.82%	69.59%	1.90%	27.69%
5:	10000	TRUE	power	TRUE	TRUE	10	50.50%	2.92%	39.60%	6.98%
6:	10000	TRUE	typeI	TRUE	TRUE	10	0.66%	70.13%	1.90%	27.31%
7:	10000	TRUE	power	TRUE	TRUE	5	48.22%	3.19%	41.57%	7.02%
8:	10000	TRUE	typeI	TRUE	TRUE	5	0.69%	68.11%	1.93%	29.27%
9:	9900	TRUE	power	FALSE	TRUE	10	50.46%	3.36%	39.87%	6.30%
10:	10000	TRUE	typeI	FALSE	TRUE	10	0.44%	0.23%	1.71%	97.62%
11:	9900	TRUE	power	FALSE	TRUE	5	48.22%	3.18%	42.25%	6.34%
12:	10000	TRUE	typeI	FALSE	TRUE	5	0.55%	0.09%	1.75%	97.61%
13:	9900	TRUE	power	FALSE	FALSE	10	51.20%	3.32%	39.46%	6.01%
14:	10000	TRUE	typeI	FALSE	FALSE	10	0.58%	0.10%	1.69%	97.63%
15:	9900	TRUE	power	FALSE	FALSE	5	48.71%	3.14%	41.95%	6.20%
16:	10000	TRUE	typeI	FALSE	FALSE	5	0.62%	0.02%	1.74%	97.62%
17:	10000	FALSE	power	TRUE	FALSE	5	47.04%	2.46%	43.34%	7.16%
18:	9900	FALSE	typeI	TRUE	FALSE	5	0.82%	67.97%	1.88%	29.33%

Method 3:

	N	missing	hypo	binding	fixC	ar	${\tt decision.eff}$	${\tt decision.fut}$	final.eff	final.fut
1:	10000	TRUE	power	TRUE	FALSE	10	54.76%	3.19%	35.42%	6.63%
2:	10000	TRUE	typeI	TRUE	FALSE	10	0.88%	68.45%	1.80%	28.87%
3:	10000	TRUE	power	TRUE	FALSE	5	49.73%	3.32%	40.05%	6.90%
4:	10000	TRUE	typeI	TRUE	FALSE	5	0.80%	67.87%	1.90%	29.43%
5:	10000	TRUE	power	TRUE	TRUE	10	54.76%	3.19%	35.42%	6.63%
6:	10000	TRUE	typeI	TRUE	TRUE	10	0.88%	68.45%	1.80%	28.87%
7:	10000	TRUE	power	TRUE	TRUE	5	49.73%	3.32%	40.05%	6.90%
8:	10000	TRUE	typeI	TRUE	TRUE	5	0.80%	67.87%	1.90%	29.43%
9:	9900	TRUE	power	FALSE	TRUE	10	54.39%	3.44%	36.02%	6.14%
10:	10000	TRUE	typeI	FALSE	TRUE	10	0.61%	0.36%	1.72%	97.31%
11:	9900	TRUE	power	FALSE	TRUE	5	49.62%	3.29%	40.85%	6.24%
12:	10000	TRUE	typeI	FALSE	TRUE	5	0.59%	0.13%	1.75%	97.53%
13:	9900	TRUE	power	FALSE	FALSE	10	54.39%	3.44%	36.02%	6.14%
14:	10000	TRUE	typeI	FALSE	FALSE	10	0.61%	0.36%	1.72%	97.31%
15:	9900	TRUE	power	FALSE	FALSE	5	49.62%	3.29%	40.85%	6.24%
16:	10000	TRUE	typeI	FALSE	FALSE	5	0.59%	0.13%	1.75%	97.53%
17:	10000	FALSE	power	TRUE	FALSE	5	47.93%	2.75%	42.16%	7.16%
18:	9900	FALSE	typeI	TRUE	FALSE	5	0.77%	66.12%	1.89%	31.22%

Relative frequency of stopping for with a threshold below 1.96:

	scenario	${\tt missing}$	${\tt method}$	binding	fixC	ar	hypo	N	${\tt rejection}$	rejectionBelow196
1:	1	TRUE	1	TRUE	FALSE	10	power	10000	90.43%	0.43%
2:	1	TRUE	2	TRUE	FALSE	10	power	10000	90.37%	0.47%
3:	2	TRUE	1	TRUE	FALSE	10	typeI	10000	2.76%	0.20%
4:	2	TRUE	2	TRUE	FALSE	10	typeI	10000	2.75%	0.20%
5:	3	TRUE	1	TRUE	FALSE	5	power	10000	90.13%	0.40%
6:	3	TRUE	2	TRUE	FALSE	5	power	10000	90.11%	0.40%
7:	4	TRUE	1	TRUE	FALSE	5	typeI	10000	2.72%	0.10%
8:	4	TRUE	2	TRUE	FALSE	5	typeI	10000	2.72%	0.10%
9:	13	TRUE	1	FALSE	FALSE	10	power	9900	90.74%	0.54%
10:	13	TRUE	2	FALSE	FALSE	10	power	9900	90.67%	0.56%
11:	14	TRUE	1	FALSE	FALSE	10	typeI	10000	2.29%	0.15%
12:	14	TRUE	2	FALSE	FALSE	10	typeI	10000	2.27%	0.13%
13:	15	TRUE	1	FALSE	FALSE	5	power	9900	90.67%	0.35%
14:	15	TRUE	2	FALSE	FALSE	5	power	9900	90.66%	0.36%
15:	16	TRUE	1	FALSE	FALSE	5	typeI	10000	2.36%	0.07%
16:	16	TRUE	2	FALSE	FALSE	5	typeI	10000	2.36%	0.07%
17:	17	FALSE	1	TRUE	FALSE	5	power	10000	90.38%	0.43%
18:	17	FALSE	2	TRUE	FALSE	5	power	10000	90.38%	0.43%
19:	18	FALSE	1	TRUE	FALSE	5	typeI	9900	2.70%	0.10%
20:	18	FALSE	2	TRUE	FALSE	5	typeI	9900	2.70%	0.10%

2.2 3 stages

Relative frequency of stopping for efficacy/futility at decision/final

• Method 1

	N	missing	hypo	binding	fixC	ar	dec1.eff	dec1.fut	dec2.eff	dec2.fut	final.eff	final.fut
1:	10000	TRUE	power	TRUE	FALSE	10	28.83%	1.34%	34.78%	2.69%	27.35%	5.01%
2:	10000	TRUE	typeI	TRUE	FALSE	10	0.47%	47.36%	0.63%	35.07%	1.36%	15.11%
3:	10000	TRUE	power	TRUE	FALSE	5	26.51%	1.38%	34.74%	2.85%	29.37%	5.15%
4:	10000	TRUE	typeI	TRUE	FALSE	5	0.44%	44.88%	0.69%	35.90%	1.39%	16.70%
5:	10000	TRUE	power	TRUE	TRUE	10	28.46%	1.71%	34.49%	2.98%	27.35%	5.01%
6:	10000	TRUE	typeI	TRUE	TRUE	10	0.34%	47.49%	0.55%	35.15%	1.36%	15.11%
7:	10000	TRUE	power	TRUE	TRUE	5	26.31%	1.58%	34.53%	3.06%	29.37%	5.15%
8:	10000	TRUE	typeI	TRUE	TRUE	5	0.41%	44.91%	0.61%	35.98%	1.39%	16.70%
9:	10000	TRUE	power	FALSE	TRUE	10	28.18%	1.89%	35.13%	2.93%	26.70%	5.17%
10:	9931	TRUE	typeI	FALSE	TRUE	10	0.35%	0.14%	0.49%	0.40%	1.45%	97.16%
11:	10000	TRUE	power	FALSE	TRUE	5	26.02%	1.75%	35.41%	2.97%	28.51%	5.34%
12:	10000	TRUE	typeI	FALSE	TRUE	5	0.42%	0.01%	0.58%	0.08%	1.50%	97.41%
13:	10000	TRUE	power	FALSE	FALSE	10	28.52%	1.55%	35.38%	2.68%	26.70%	5.17%
14:	9931	TRUE	typeI	FALSE	FALSE	10	0.44%	0.05%	0.55%	0.34%	1.45%	97.16%
15:	10000	TRUE	power	FALSE	FALSE	5	26.21%	1.56%	35.59%	2.79%	28.51%	5.34%
16:	10000	TRUE	typeI	FALSE	FALSE	5	0.43%	0	0.59%	0.07%	1.50%	97.41%
17:	10000	FALSE	power	TRUE	FALSE	5	25.38%	1.48%	34.27%	2.45%	30.20%	6.22%
18:	10000	FALSE	typeI	TRUE	FALSE	5	0.35%	42.84%	0.67%	36.11%	1.63%	18.40%

• Method 2

	N	missing	hypo	binding	fixC	ar	dec1.eff	dec1.fut	dec2.eff	dec2.fut	final.eff	final.fut
1:	10000	TRUE	power	TRUE	FALSE	10	28.87%	1.42%	34.79%	2.82%	27.19%	4.91%
2:	10000	TRUE	typeI	TRUE	FALSE	10	0.47%	47.79%	0.65%	35.40%	1.39%	14.30%
3:	10000	TRUE	power	TRUE	FALSE	5	26.53%	1.38%	34.80%	2.86%	29.28%	5.15%
4:	10000	TRUE	typeI	TRUE	FALSE	5	0.44%	44.92%	0.69%	35.99%	1.39%	16.57%
5:	10000	TRUE	power	TRUE	TRUE	10	28.18%	1.51%	34.40%	2.85%	27.86%	5.20%
6:	10000	TRUE	typeI	TRUE	TRUE	10	0.33%	44.96%	0.53%	36.43%	1.38%	16.37%
7:	10000	TRUE	power	TRUE	TRUE	5	26.18%	1.41%	34.45%	2.84%	29.71%	5.41%
8:	10000	TRUE	typeI	TRUE	TRUE	5	0.41%	43.53%	0.59%	36.40%	1.43%	17.64%
9:	10000	TRUE	power	FALSE	TRUE	10	28.03%	1.62%	35.14%	2.82%	27.10%	5.29%
10:	9931	TRUE	typeI	FALSE	TRUE	10	0.35%	0.12%	0.48%	0.35%	1.48%	97.21%
11:	10000	TRUE	power	FALSE	TRUE	5	25.81%	1.60%	35.45%	2.72%	28.86%	5.56%
12:	10000	TRUE	typeI	FALSE	TRUE	5	0.40%	0.01%	0.59%	0.07%	1.52%	97.41%
13:	10000	TRUE	power	FALSE	FALSE	10	28.53%	1.61%	35.47%	2.88%	26.49%	5.02%
14:	9931	TRUE	typeI	FALSE	FALSE	10	0.44%	0.05%	0.52%	0.40%	1.45%	97.13%
15:	10000	TRUE	power	FALSE	FALSE	5	26.21%	1.56%	35.66%	2.81%	28.44%	5.32%

16:	10000	TRUE typeI	FALSE FALSE	5	0.43%	0	0.60%	0.07%	1.49%	97.41%
17:	10000	FALSE power	TRUE FALSE	5	25.42%	1.50%	34.24%	2.50%	30.15%	6.19%
18:	10000	FALSE typeI	TRUE FALSE	5	0.35%	42.87%	0.67%	36.23%	1.63%	18.25%

• Method 3

	N	missing	hypo	binding	fixC	ar	dec1.eff	dec1.fut	dec2.eff	dec2.fut	final.eff	final.fut
1:	10000	TRUE	power	TRUE	FALSE	10	31.44%	1.68%	35.60%	2.80%	23.40%	5.08%
2:	10000	TRUE	typeI	TRUE	FALSE	10	0.42%	43.69%	0.61%	36.37%	1.41%	17.50%
3:	10000	TRUE	power	TRUE	FALSE	5	26.95%	1.47%	35.22%	2.96%	28.11%	5.29%
4:	10000	TRUE	typeI	TRUE	FALSE	5	0.41%	43.52%	0.62%	36.31%	1.46%	17.68%
5:	10000	TRUE	power	TRUE	TRUE	10	31.44%	1.68%	35.60%	2.80%	23.40%	5.08%
6:	10000	TRUE	typeI	TRUE	TRUE	10	0.42%	43.69%	0.61%	36.37%	1.41%	17.50%
7:	10000	TRUE	power	TRUE	TRUE	5	26.95%	1.47%	35.22%	2.96%	28.11%	5.29%
8:	10000	TRUE	typeI	TRUE	TRUE	5	0.41%	43.52%	0.62%	36.31%	1.46%	17.68%
9:	10000	TRUE	power	FALSE	TRUE	10	30.84%	1.66%	36.22%	2.90%	23.34%	5.04%
10:	9931	TRUE	typeI	FALSE	TRUE	10	0.41%	0.16%	0.63%	0.58%	1.42%	96.79%
11:	10000	TRUE	power	FALSE	TRUE	5	26.83%	1.67%	35.85%	2.85%	27.36%	5.44%
12:	10000	TRUE	typeI	FALSE	TRUE	5	0.45%	0.02%	0.67%	0.13%	1.52%	97.21%
13:	10000	TRUE	power	FALSE	FALSE	10	30.84%	1.66%	36.22%	2.90%	23.34%	5.04%
14:	9931	TRUE	typeI	FALSE	FALSE	10	0.41%	0.16%	0.63%	0.58%	1.42%	96.79%
15:	10000	TRUE	power	FALSE	FALSE	5	26.83%	1.67%	35.85%	2.85%	27.36%	5.44%
16:	10000	TRUE	typeI	FALSE	FALSE	5	0.45%	0.02%	0.67%	0.13%	1.52%	97.21%
17:	10000	FALSE	power	TRUE	FALSE	5	25.94%	1.64%	34.58%	2.53%	29.05%	6.26%
18:	10000	FALSE	typeI	TRUE	FALSE	5	0.32%	41.33%	0.62%	36.51%	1.58%	19.64%

Relative frequency of stopping for with a threshold below 1.96:

	scenario	missing	method	binding	fixC	ar	hypo	N	rejection	rejectionBelow196
1:	1	TRUE	1	TRUE	FALSE	10	power	10000	90.96%	0.66%
2:	1	TRUE	2	TRUE	FALSE	10	power	10000	90.85%	0.62%
3:	2	TRUE	1	TRUE	FALSE	10	typeI	10000	2.46%	0.21%
4:	2	TRUE	2	TRUE	FALSE	10	typeI	10000	2.51%	0.21%
5:	3	TRUE	1	TRUE	FALSE	5	power	10000	90.62%	0.41%
6:	3	TRUE	2	TRUE	FALSE	5	power	10000	90.61%	0.42%
7:	4	TRUE	1	TRUE	FALSE	5	typeI	10000	2.52%	0.11%
8:	4	TRUE	2	TRUE	FALSE	5	typeI	10000	2.52%	0.11%
9:	13	TRUE	1	FALSE	FALSE	10	power	10000	90.60%	0.59%
10:	13	TRUE	2	FALSE	FALSE	10	power	10000	90.49%	0.57%
11:	14	TRUE	1	FALSE	FALSE	10	typeI	9931	2.45%	0.15%
12:	14	TRUE	2	FALSE	FALSE	10	typeI	9931	2.42%	0.13%
13:	15	TRUE	1	FALSE	FALSE	5	power	10000	90.31%	0.37%
14:	15	TRUE	2	FALSE	FALSE	5	power	10000	90.31%	0.38%
15:	16	TRUE	1	FALSE	FALSE	5	typeI	10000	2.52%	0.02%
16:	16	TRUE	2	FALSE	FALSE	5	typeI	10000	2.52%	0.02%
17:	17	FALSE	1	TRUE	FALSE	5	power	10000	89.85%	0.39%
18:	17	FALSE	2	TRUE	FALSE	5	power	10000	89.81%	0.38%
19:	18	FALSE	1	TRUE	FALSE	5	typeI	10000	2.65%	0.13%
20:	18	FALSE	2	TRUE	FALSE	5	typeI	10000	2.65%	0.13%

3 Bias (True effect: 0.6 under the alternative)

3.1 2 stages

Bias per estimator and method¹:

```
hypo missing binding fixC ar biasMLE1 biasMLE2 biasMLE3 biasMUE1 biasMUE2 biasMUE3
                    TRUE FALSE 10 0.01251 0.01233 0.01275 0.00470 0.00452 -0.00450
1: power
            TRUE
2: typeI
                    TRUE FALSE 10 -0.01673 -0.01655 -0.01713 -0.00518 -0.00508 -0.01539
            TRUE
                    TRUE FALSE 5 0.02412 0.02410 0.02442 0.01141 0.01138 0.00302
3: power
            TRUE
4: typeI
            TRUE
                    TRUE FALSE 5 -0.02839 -0.02837 -0.02897 -0.01206 -0.01207 -0.02288
5: power
            TRUE
                    TRUE
                          TRUE 10 0.01251 0.01278 0.01275 -0.01829 -0.01939 -0.00450
                          TRUE 10 -0.01673 -0.01725 -0.01713 -0.04244 -0.04509 -0.01539
6: typeI
            TRUE
                    TRUE
7: power
            TRUE
                    TRUE
                          TRUE
                                  0.02412 0.02444 0.02442 -0.02060 -0.02149 0.00302
                          TRUE 5 -0.02839 -0.02907 -0.02897 -0.06515 -0.06704 -0.02288
8: typeI
            TRUE
                    TRUE
            TRUE
                   FALSE
                          TRUE 10 0.01191 0.01234 0.01241 -0.00291 -0.00536
9: power
                                                                               0.01012
10: typeI
            TRUE
                   FALSE
                          TRUE 10 -0.00049 -0.00049 -0.00029 -0.00156 -0.00154
                                                                                0.00000
11: power
                                  0.02382 0.02410 0.02409 -0.00384 -0.00573
            TRUE
                   FALSE
                          TRUE
                                                                                0.01884
12: typeI
            TRUE
                   FALSE
                          TRUE
                               5 -0.00004 -0.00004 0.00004 -0.00057 -0.00056 -0.00010
13: power
            TRUE
                   FALSE FALSE 10 0.01191 0.01174 0.01241 0.01785 0.01836
                                                                                0.01012
14: typeI
            TRUE
                   FALSE FALSE 10 -0.00049 -0.00049 -0.00029 -0.00060 -0.00072
                                                                                0.00000
15: power
            TRUE
                   FALSE FALSE
                               5
                                  0.02382 0.02379
                                                    0.02409 0.02577 0.02582
                                                                                0.01884
16: typeI
            TRUE
                   FALSE FALSE
                                5 -0.00004 -0.00004 0.00004 -0.00002 -0.00002 -0.00010
17: power
           FALSE
                    TRUE FALSE
                               5 0.02542 0.02542 0.02598 0.01365 0.01366 0.00371
18: typeI
           FALSE
                    TRUE FALSE 5 -0.02471 -0.02472 -0.02520 -0.00941 -0.00941 -0.02205
```

Median bias ² per estimator and method:

	hypo	missing	binding	fixC	ar	${\tt mbiasMLE1}$	${\tt mbiasMLE2}$	${\tt mbiasMLE3}$	${\tt mbiasMUE1}$	${\tt mbias MUE2}$	mbiasMUE3
1:	power	TRUE	TRUE	FALSE	10	0.0218	0.0218	0.0198	-0.00560	-0.00550	-0.01340
2:	typeI	TRUE	TRUE	FALSE	10	-0.0272	-0.0265	-0.0296	-0.00520	-0.00606	-0.02692
3:	power	TRUE	TRUE	FALSE	5	0.0472	0.0472	0.0471	-0.00425	-0.00425	-0.00760
4:	typeI	TRUE	TRUE	FALSE	5	-0.0446	-0.0444	-0.0478	-0.00700	-0.00746	-0.02836
5:	power	TRUE	TRUE	TRUE	10	0.0218	0.0216	0.0198	-0.02751	-0.02951	-0.01340
6:	typeI	TRUE	TRUE	TRUE	10	-0.0272	-0.0292	-0.0296	-0.08467	-0.08715	-0.02692
7:	power	TRUE	TRUE	TRUE	5	0.0472	0.0471	0.0471	-0.02956	-0.03136	-0.00760
8:	typeI	TRUE	TRUE	TRUE	5	-0.0446	-0.0472	-0.0478	-0.10726	-0.10396	-0.02836
9:	power	TRUE	FALSE	TRUE	10	0.0256	0.0257	0.0261	-0.00808	-0.01103	0.00434
10:	typeI	TRUE	FALSE	TRUE	10	-0.0040	-0.0040	-0.0040	-0.00516	-0.00516	-0.00301
11:	power	TRUE	FALSE	TRUE	5	0.0499	0.0495	0.0503	-0.01402	-0.01674	0.00559
12:	typeI	TRUE	FALSE	TRUE	5	-0.0040	-0.0040	-0.0040	-0.00455	-0.00455	-0.00396
13:	power	TRUE	FALSE	FALSE	10	0.0256	0.0256	0.0261	0.01372	0.01458	0.00434
14:	typeI	TRUE	FALSE	FALSE	10	-0.0040	-0.0040	-0.0040	-0.00440	-0.00450	-0.00301
15:	power	TRUE	FALSE	FALSE	5	0.0499	0.0499	0.0503	0.00871	0.00881	0.00559

¹e.g. biasMLE1 mixed model estimator (treatment effect), method 1 (boundaries)

²Relative frequency at which the estimate is greater than the truth minus 0.5

16: typeI FALSE FALSE 5 -0.0040 -0.0040 -0.00420 -0.00420 -0.00396 TRUE -0.0040 17: power TRUE FALSE 5 0.0479 0.0479 0.0481 -0.00425 -0.00435 -0.01240 FALSE 18: typeI TRUE FALSE 5 -0.0417 -0.0418 -0.0437 -0.00425 -0.00445 -0.02922 FALSE

3.2 3 stages

Bias per estimator and method³:

	hypo	missing	binding	fixC	ar	biasMLE1	biasMLE2	biasMLE3	biasMUE1	biasMUE2	biasMUE3
1:	power	TRUE	TRUE	FALSE	10	0.0240	0.0238	0.0255	0.0161	0.0159	0.0074
2:	typeI	TRUE	TRUE	FALSE	10	-0.0281	-0.0279	-0.0280	-0.0134	-0.0129	-0.0249
3:	power	TRUE	TRUE	FALSE	5	0.0380	0.0380	0.0388	0.0216	0.0217	0.0130
4:	typeI	TRUE	TRUE	FALSE	5	-0.0447	-0.0447	-0.0448	-0.0226	-0.0226	-0.0348
5:	power	TRUE	TRUE	TRUE	10	0.0240	0.0245	0.0255	-0.0076	-0.0083	0.0074
6:	typeI	TRUE	TRUE	TRUE	10	-0.0281	-0.0283	-0.0280	-0.0562	-0.0582	-0.0249
7:	power	TRUE	TRUE	TRUE	5	0.0380	0.0386	0.0388	-0.0105	-0.0104	0.0130
8:	typeI	TRUE	TRUE	TRUE	5	-0.0447	-0.0449	-0.0448	-0.0848	-0.0859	-0.0348
9:	power	TRUE	FALSE	TRUE	10	0.0242	0.0246	0.0253	0.0150	0.0129	0.0274
10:	typeI	TRUE	FALSE	TRUE	10	0.0024	0.0023	0.0027	0.0010	0.0011	0.0031
11:	power	TRUE	FALSE	TRUE	5	0.0380	0.0385	0.0388	0.0133	0.0112	0.0353
12:	typeI	TRUE	FALSE	TRUE	5	0.0035	0.0035	0.0038	0.0029	0.0029	0.0038
13:	power	TRUE	FALSE	FALSE	10	0.0242	0.0241	0.0253	0.0348	0.0356	0.0274
14:	typeI	TRUE	FALSE	FALSE	10	0.0024	0.0024	0.0027	0.0021	0.0019	0.0031
15:	power	TRUE	FALSE	FALSE	5	0.0380	0.0380	0.0388	0.0427	0.0428	0.0353
16:	typeI	TRUE	FALSE	FALSE	5	0.0035	0.0035	0.0038	0.0032	0.0033	0.0038
17:	power	FALSE	TRUE	FALSE	5	0.0373	0.0372	0.0379	0.0221	0.0220	0.0122
18:	typeI	FALSE	TRUE	FALSE	5	-0.0420	-0.0419	-0.0423	-0.0203	-0.0202	-0.0344

Median bias 4 per estimator and method:

	hypo	missing	${\tt binding}$	fixC	ar	${\tt mbiasMLE1}$	${\tt mbiasMLE2}$	${\tt mbiasMLE3}$	mbiasMUE1	${\tt mbias MUE2}$	mbiasMUE3
1:	power	TRUE	TRUE	FALSE	10	0.0348	0.0346	0.0360	0.00130	0.00120	-0.0128
2:	typeI	TRUE	TRUE	FALSE	10	-0.0428	-0.0423	-0.0426	0.00205	0.00200	-0.0161
3:	power	TRUE	TRUE	FALSE	5	0.0577	0.0576	0.0573	-0.00085	-0.00085	-0.0185
4:	typeI	TRUE	TRUE	FALSE	5	-0.0589	-0.0588	-0.0589	-0.00455	-0.00415	-0.0195
5:	power	TRUE	TRUE	TRUE	10	0.0348	0.0350	0.0360	-0.03457	-0.03658	-0.0128
6:	typeI	TRUE	TRUE	TRUE	10	-0.0428	-0.0426	-0.0426	-0.02798	-0.03293	-0.0161
7:	power	TRUE	TRUE	TRUE	5	0.0577	0.0578	0.0573	-0.05581	-0.05901	-0.0185
8:	typeI	TRUE	TRUE	TRUE	5	-0.0589	-0.0589	-0.0589	-0.04987	-0.05342	-0.0192
9:	power	TRUE	FALSE	TRUE	10	0.0413	0.0408	0.0420	-0.00557	-0.01015	0.0046
10:	typeI	TRUE	FALSE	TRUE	10	0.0012	0.0012	0.0012	-0.00101	-0.00066	0.0026
11:	power	TRUE	FALSE	TRUE	5	0.0633	0.0633	0.0629	-0.03128	-0.03685	0.0062
12:	typeI	TRUE	FALSE	TRUE	5	0.0015	0.0015	0.0015	0.00075	0.00070	0.0023
13:	power	TRUE	FALSE	FALSE	10	0.0413	0.0415	0.0420	0.02261	0.02398	0.0046
14:	typeI	TRUE	FALSE	FALSE	10	0.0012	0.0012	0.0012	-0.00025	-0.00010	0.0026
15:	power	TRUE	FALSE	FALSE	5	0.0633	0.0630	0.0629	0.02410	0.02421	0.0063
16:	typeI	TRUE	FALSE	FALSE	5	0.0015	0.0015	0.0015	0.00085	0.00075	0.0022
17:	power	FALSE	TRUE	FALSE	5	0.0584	0.0586	0.0585	0.00285	0.00275	-0.0157

 $^{^3}$ e.g. biasMLE1 mixed model estimator (treatment effect), method 1 (boundaries)

 $^{^4}$ Relative frequency at which the estimate is greater than the truth minus 0.5

18: typeI FALSE TRUE FALSE 5 -0.0589 -0.0584 -0.0587 0.00190 0.00230 -0.0117

4 Distribution of the estimates

4.1 2 stages

Distribution of the estimates:

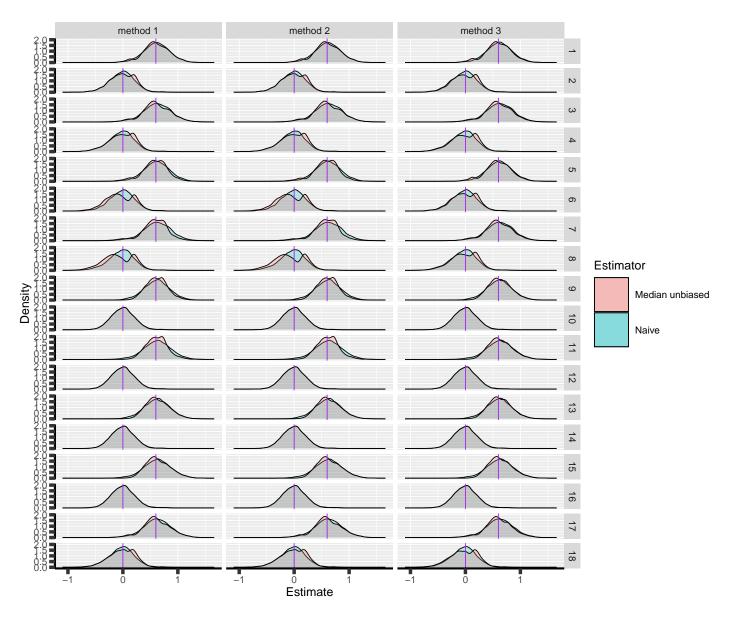


Figure 5: Naive and Median unbiased estimate distribution over all simulations. Each row correspond to a different scenario

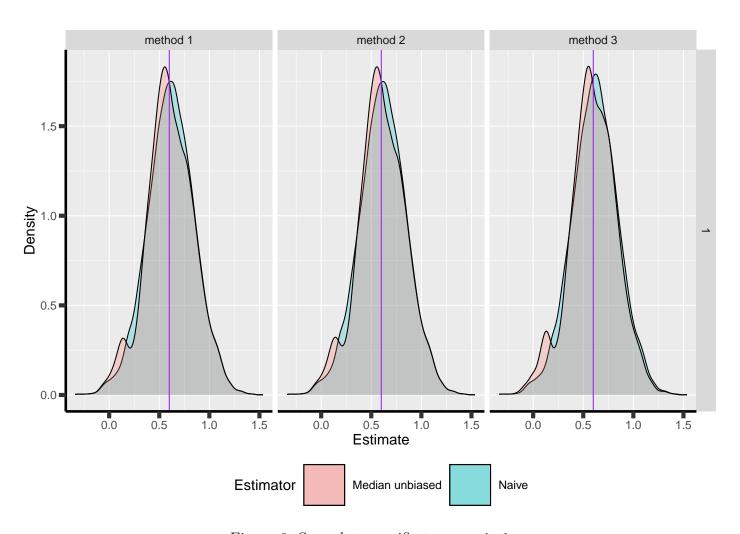


Figure 6: Same but specific to scenario 1

Distribution of the median unbiased estimate conditional to the stage:

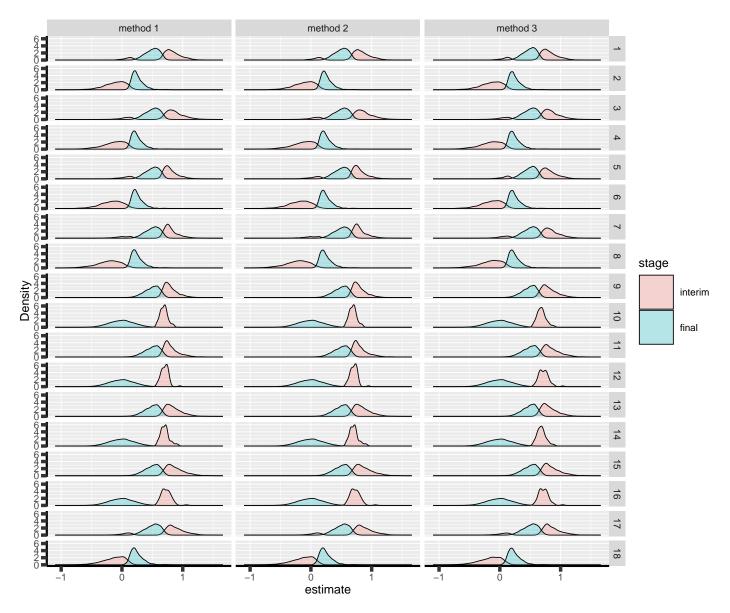


Figure 7: Median unbiased estimate distribution conditional to the stage. Each row correspond to a different scenario.

4.2 3 stages

Distribution of the estimates:

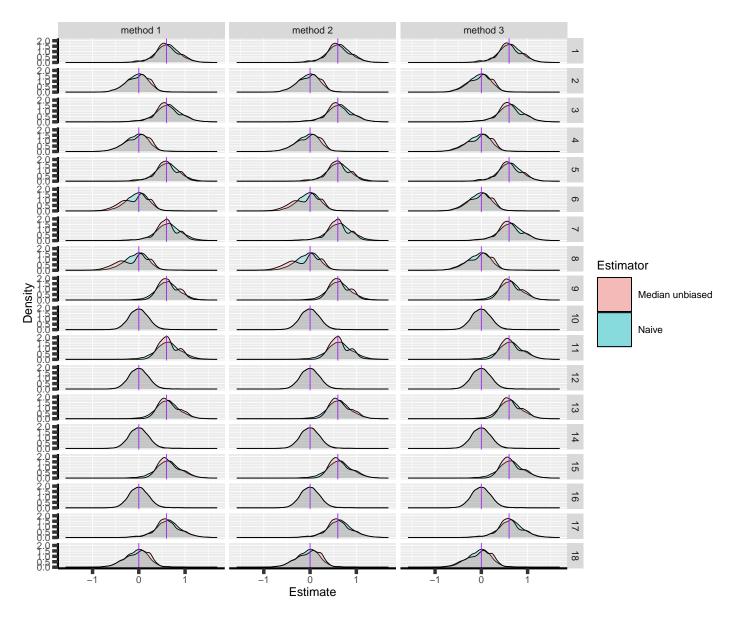


Figure 8: Naive and Median unbiased estimate distribution over all simulations. Each row correspond to a different scenario

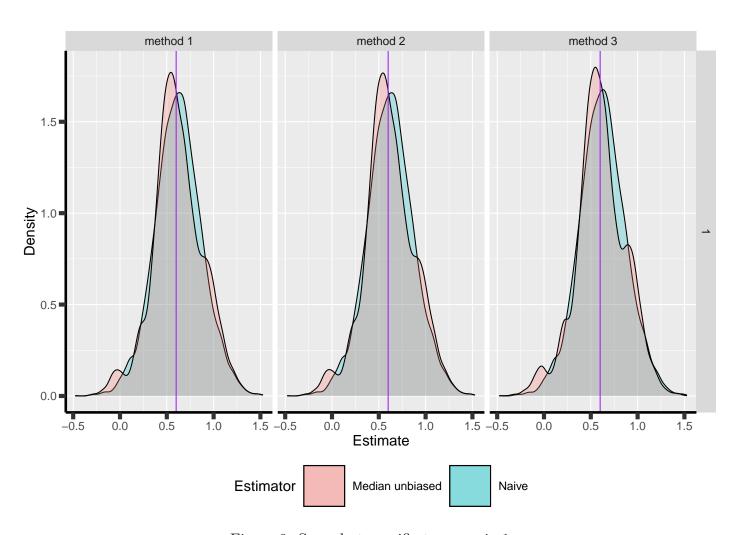


Figure 9: Same but specific to scenario 1

Distribution of the median unbiased estimate conditional to the stage:

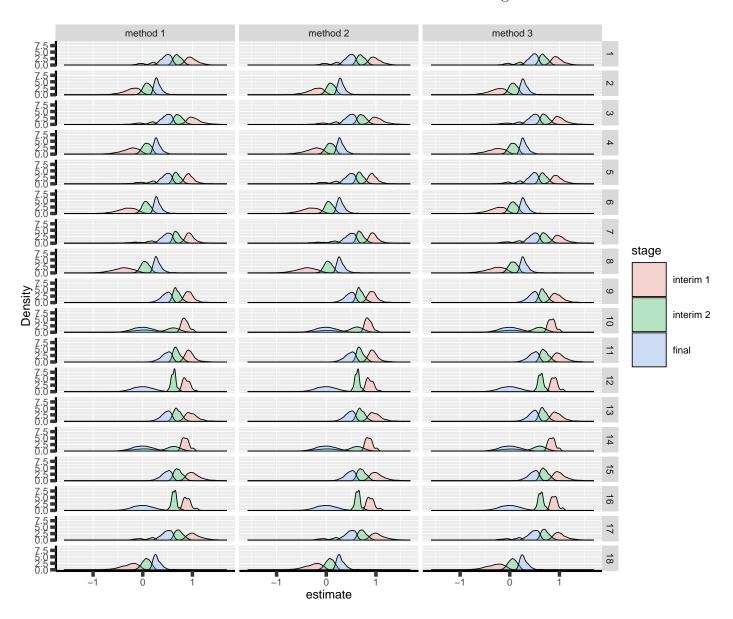


Figure 10: Median unbiased estimate distribution conditional to the stage. Each row correspond to a different scenario.

5 Special cases

5.1 2 stages

Reason for stopping (efficacy, futility, Imax reached), continuing the trial (decreasing information, no boundary crossed), or concluding (stop for futility at interim):

		scenario	1	2	3	4	5	6	7	8		
reason	method											
efficacy	1		5093	86	4874	80	5093	86	4874	80		
	2		5093	86	4874	81	5068	86	4861	77		
	3		5553	128	5028	91	5553	128	5028	91		
futility	1		320	7179	299	6950	320	7179	299	6950		
	2		349	7269	301	6960	274	6993	280	6803		
	3		242	6805	277	6776	242	6805	277	6776		
no boundary crossed	1		4587	2735	4827	2970	4587	2735	4827	2970		
	2		4558	2645	4825	2959	4658	2921	4859	3120		
	3		4205	3067	4695	3133	4205	3067	4695	3133		
stop for futility at interim	1		0	0	0	0	0	0	0	0		
	2		0	0	0	0	0	0	0	0		
	3		10	3	0	0	10	3	0	0		
		scenario	9	10	11	12	13	14	15	16	17	18
reason	method	Scenar 10	9	10	11	12	10	14	10	10	11	10
efficacy	1		5039	68	4821	64	5039	68	4821	64	4700	82
efficacy	2		5028		4806		5039		4821		4700	82
	3		5460		4959		5460		4959		4845	90
futility	1			7164		6964		7164		6964		6719
Tuciffcy	2			6978		6833		7258		6973		6728
	3			6800		6816		6800		6816		6532
	J		200	0000	213	0010	200	0000	213	0010		
no boundary crossed	1		4523	2768	4770	2072	4523	2768	4770	2972	5050	
no boundary crossed	1									2972		
no boundary crossed	2		4571	2955	4811	3103	4502	2674	4767	2963	5050	3090
·	2 3		4571 4174	2955 3103	4811 4662	3103 3112	4502 4174	2674 3103	4767 4662	2963 3112	5050 4932	3090 3278
stop for futility at interim	2 3 1		4571 4174 0	2955 3103 0	4811 4662 0	3103 3112 0	4502 4174 0	2674 3103 0	4767 4662 0	2963 3112 0	5050 4932 0	3090 3278 0
·	2 3		4571 4174	2955 3103	4811 4662	3103 3112	4502 4174	2674 3103	4767 4662	2963 3112	5050 4932	3090 3278

5.2 3 stages

Reason for stopping (efficacy, futility, Imax reached), continuing the trial (decreasing information, no boundary crossed), or concluding (stop for futility at interim):

		scenario	1	2	3	4	5	6	7	8	
reason	method										
efficacy	1		6306	115	6116	113	6306	115	6116	113	
	2		6305	115	6123	113	6275	114	6095	111	
	3		6785	155	6270	120	6785	155	6270	120	
futility	1		444	8228	432	8078	444	8228	432	8078	
	2		469	8306	434	8091	412	8102	393	7982	
	3		357	7943	390	7966	357	7943	390	7966	
Imax reached	1		14	10	0	0	14	10	0	0	
	2		16	10	0	0	7	9	0	0	
	3		10	11	0	0	10	11	0	0	
no boundary crossed	1	1	.0219	6864	10663	7277	10219	6864	10663	7277	
	2	1	.0181	6743	10652	7260	10337	7246	10753	7513	
	3		9536	7480	10498	7521	9536	7480	10498	7521	
stop for futility at interim	1		0	0	0	0	0	0	0	0	
	2		0	0	0	0	0	0	0	0	
	3		24	2	3	0	24	2	3	0	
		scenario	9	10	11	12	13	14	15	16	17
reason	method	boonario	J	10		12	10		10	10	
efficacy	1		6334	116	6177	109	6334	116	6177	109	5961
y	2		6322	113	6160	107	6337	116	6182	110	5963
	3		6782	156	6325	127	6782	156	6325	127	6105
futility	1			12836		12525		12836		12525	397
v	2		427	12493		12326		12946		12540	403
	3			12227		12319		12227		12319	364
Imax reached	1		14	22	0	0	14	22	0	0	0
	2		12	17	0	0	16	25	0	0	0
	3		12	22	0	0	12	22	0	0	0
no boundary crossed	1	1	.0180		10608	7323	10180		10608	7323	10956
•	2		.0274		10701		10137		10599		10942
	3		9588		10430	7507			10430		10773
stop for futility at interim	1		0	0	0	0	0	0	0	0	0
-	2		0	0	0	0	0	0	0	0	0
	3		27	0	1	0	27	0	1	0	0

6 Reversal probability

6.1 2 stages

Percentage of time we observe a reversal:

	N	hypo	missing	ar	binding	fixC	fu2eff_1	fu2eff_2	fu2eff_3	eff2fu_1	eff2fu_2	eff2fu_3
1:	10000	power	TRUE	10	TRUE	FALSE	0.43%	0.50%	0	0.13%	0.14%	0.77%
2:	10000	typeI	TRUE	10	TRUE	FALSE	0.13%	0.13%	0	0.13%	0.13%	0.40%
3:	10000	power	TRUE	5	TRUE	FALSE	0.05%	0.05%	0	0.05%	0.05%	0.55%
4:	10000	typeI	TRUE	5	TRUE	FALSE	0.03%	0.03%	0	0.02%	0.02%	0.11%
5:	10000	power	TRUE	10	TRUE	TRUE	0.24%	0.19%	0	0.37%	0.37%	0.77%
6:	10000	typeI	TRUE	10	TRUE	TRUE	0.04%	0.04%	0	0.24%	0.24%	0.40%
7:	10000	power	TRUE	5	TRUE	TRUE	0	0	0	0.40%	0.39%	0.55%
8:	10000	typeI	TRUE	5	TRUE	TRUE	0	0	0	0.09%	0.08%	0.11%
9:	9900	power	TRUE	10	FALSE	TRUE	0.21%	0.17%	0	0.49%	0.49%	0.76%
10:	10000	typeI	TRUE	10	FALSE	TRUE	0	0	0	0.23%	0.23%	0.36%
11:	9900	power	TRUE	5	FALSE	TRUE	0	0	0	0.35%	0.32%	0.47%
12:	10000	typeI	TRUE	5	FALSE	TRUE	0	0	0	0.09%	0.09%	0.13%
13:	9900	power	TRUE	10	FALSE	FALSE	0.40%	0.46%	0	0.15%	0.16%	0.76%
14:	10000	typeI	TRUE	10	FALSE	FALSE	0	0	0	0.08%	0.10%	0.36%
15:	9900	power	TRUE	5	FALSE	FALSE	0.04%	0.05%	0	0.04%	0.04%	0.47%
16:	10000	typeI	TRUE	5	FALSE	FALSE	0	0	0	0.02%	0.02%	0.13%
17:	10000	power	FALSE	5	TRUE	FALSE	0.08%	0.08%	0	0.04%	0.04%	0.52%
18:	9900	typeI	FALSE	5	TRUE	FALSE	0.02%	0.02%	0	0.03%	0.03%	0.14%

6.2 3 stages

Percentage of time we observe a reversal:

	N	hypo	missing	ar	binding	fixC	fu2eff_1	fu2eff_2	${\tt fu2eff_3}$	${\tt eff2fu_1}$	${\tt eff2fu_2}$	eff2fu_3
1:	10000	power	TRUE	10	TRUE	FALSE	0.57%	0.66%	0	0.15%	0.19%	0.91%
2:	10000	typeI	TRUE	10	TRUE	FALSE	0.16%	0.18%	0	0.22%	0.22%	0.52%
3:	10000	power	TRUE	5	TRUE	FALSE	0.15%	0.16%	0	0.06%	0.06%	0.53%
4:	10000	typeI	TRUE	5	TRUE	FALSE	0.02%	0.02%	0	0.02%	0.02%	0.17%
5:	10000	power	TRUE	10	TRUE	TRUE	0.37%	0.36%	0	0.61%	0.60%	0.91%
6:	10000	typeI	TRUE	10	TRUE	TRUE	0.06%	0.04%	0	0.33%	0.32%	0.52%
7:	10000	power	TRUE	5	TRUE	TRUE	0.05%	0.03%	0	0.37%	0.35%	0.53%
8:	10000	typeI	TRUE	5	TRUE	TRUE	0.01%	0	0	0.12%	0.11%	0.17%
9:	10000	power	TRUE	10	FALSE	TRUE	0.41%	0.39%	0	0.58%	0.56%	0.88%
10:	9931	typeI	TRUE	10	FALSE	TRUE	0	0	0	0.33%	0.30%	0.53%
11:	10000	power	TRUE	5	FALSE	TRUE	0.02%	0.01%	0	0.36%	0.35%	0.57%
12:	10000	typeI	TRUE	5	FALSE	TRUE	0	0	0	0.09%	0.08%	0.15%
13:	10000	power	TRUE	10	FALSE	FALSE	0.66%	0.74%	0	0.24%	0.26%	0.88%
14:	9931	typeI	TRUE	10	FALSE	FALSE	0	0	0	0.18%	0.21%	0.53%
15:	10000	power	TRUE	5	FALSE	FALSE	0.10%	0.12%	0	0.07%	0.07%	0.57%
16:	10000	typeI	TRUE	5	FALSE	FALSE	0	0	0	0.07%	0.07%	0.15%
17:	10000	power	FALSE	5	TRUE	FALSE	0.11%	0.10%	0	0.07%	0.07%	0.53%
18:	10000	typeI	FALSE	5	TRUE	FALSE	0.04%	0.04%	0	0.03%	0.03%	0.16%

7 Logical consistency of p-values/CIs

7.1 Mismatch p-value / boundaries

7.1.1 2 stages

When concluding for futility:

	hypo	missing	ar	binding	fixC	method	1	${\tt method}$	2	method	3
1:	power	TRUE	10	TRUE	FALSE		0		0		0
2:	typeI	TRUE	10	TRUE	FALSE		0		0		0
3:	power	TRUE	5	TRUE	FALSE		0		0		0
4:	typeI	TRUE	5	TRUE	FALSE		0		0		0
5:	power	TRUE	10	TRUE	TRUE		0		0		0
6:	typeI	TRUE	10	TRUE	TRUE		0		0		0
7:	power	TRUE	5	TRUE	TRUE		0		0		0
8:	typeI	TRUE	5	TRUE	TRUE		0		0		0
9:	power	TRUE	10	FALSE	TRUE		0		0		0
10:	typeI	TRUE	10	FALSE	TRUE		0		0		0
11:	power	TRUE	5	FALSE	TRUE		0		0		0
12:	typeI	TRUE	5	FALSE	TRUE		0		0		0
13:	power	TRUE	10	FALSE	FALSE		0		0		0
14:	typeI	TRUE	10	FALSE	FALSE		0		0		0
15:	power	TRUE	5	FALSE	FALSE		0		0		0
16:	typeI	TRUE	5	FALSE	FALSE		0		0		0
17:	power	FALSE	5	TRUE	FALSE		0		0		0
18:	typeI	FALSE	5	TRUE	FALSE		0		0		0

When concluding for efficacy:

	hypo	missing	ar	binding	fixC	${\tt method}$	1	${\tt method}$	2	${\tt method}$	3
1:	power	TRUE	10	TRUE	FALSE		0		0		0
2:	typeI	TRUE	10	TRUE	FALSE		0		0		0
3:	power	TRUE	5	TRUE	FALSE		0		0		0
4:	typeI	TRUE	5	TRUE	FALSE		0		0		0
5:	power	TRUE	10	TRUE	TRUE		0		0		0
6:	typeI	TRUE	10	TRUE	TRUE		0		0		0
7:	power	TRUE	5	TRUE	TRUE		0		0		0
8:	typeI	TRUE	5	TRUE	TRUE		0		0		0
9:	power	TRUE	10	FALSE	TRUE		0		0		0
10:	typeI	TRUE	10	FALSE	TRUE		0		0		0
11:	power	TRUE	5	FALSE	TRUE		0		0		0
12:	typeI	TRUE	5	FALSE	TRUE		0		0		0
13:	power	TRUE	10	FALSE	FALSE		0		0		0
14:	tvpeT	TRUE	10	FALSE	FALSE		0		0		0

15:	power	TRUE	5	FALSE	FALSE	0	0	0
16:	typeI	TRUE	5	FALSE	FALSE	0	0	0
17:	power	FALSE	5	TRUE	FALSE	0	0	0
18:	tvpeI	FALSE	5	TRUE	FALSE	0	0	0

7.1.2 3 stages

When concluding for futility:

	hypo	missing	ar	binding	fixC	method	1	$\tt method\ 2$	${\tt method}\ 3$
1:	power	TRUE	10	TRUE	FALSE		0	0	0
2:	typeI	TRUE	10	TRUE	FALSE		0	0	0
3:	power	TRUE	5	TRUE	FALSE		0	0	0.10%
4:	typeI	TRUE	5	TRUE	FALSE		0	0.01%	0
5:	power	TRUE	10	TRUE	TRUE		0	0	0
6:	typeI	TRUE	10	TRUE	TRUE		0	0	0
7:	power	TRUE	5	TRUE	TRUE		0	0	0.10%
8:	typeI	TRUE	5	TRUE	TRUE		0	0	0
9:	power	TRUE	10	FALSE	TRUE		0	0	0
10:	typeI	TRUE	10	FALSE	TRUE		0	0	0
11:	power	TRUE	5	FALSE	TRUE	0.10)%	0	0
12:	typeI	TRUE	5	FALSE	TRUE		0	0	0
13:	power	TRUE	10	FALSE	FALSE		0	0	0
14:	typeI	TRUE	10	FALSE	FALSE		0	0	0
15:	power	TRUE	5	FALSE	FALSE	0.10)%	0	0
16:	typeI	TRUE	5	FALSE	FALSE		0	0	0
17:	power	FALSE	5	TRUE	FALSE		0	0	0
18:	typeI	FALSE	5	TRUE	FALSE		0	0	0

Largest mismatch:

[1] 0.02499104799

When concluding for efficacy:

	hypo	missing	ar	binding	fixC	method 1	method	2	${\tt method}$	3
1:	power	TRUE	10	TRUE	FALSE	0.01%))	0		0
2:	typeI	TRUE	10	TRUE	FALSE	C)	0		0
3:	power	TRUE	5	TRUE	FALSE	C)	0		0
4:	typeI	TRUE	5	TRUE	FALSE	C)	0		0
5:	power	TRUE	10	TRUE	TRUE	0.01%	1	0		0
6:	typeI	TRUE	10	TRUE	TRUE	C)	0		0
7:	power	TRUE	5	TRUE	TRUE	C)	0		0
8:	typeI	TRUE	5	TRUE	TRUE	C)	0		0
9:	power	TRUE	10	FALSE	TRUE	C)	0		0
10:	typeI	TRUE	10	FALSE	TRUE	C)	0		0
11:	power	TRUE	5	FALSE	TRUE	C)	0		0
12:	typeI	TRUE	5	FALSE	TRUE	C)	0		0
13:	power	TRUE	10	FALSE	FALSE	C)	0	0.01	L%
14:	typeI	TRUE	10	FALSE	FALSE	C)	0		0
15:	power	TRUE	5	FALSE	FALSE	C)	0		0

16: typeI	TRUE	5	FALSE FALSE	0	0	0
17: power	FALSE	5	TRUE FALSE	0	0	0
18: typeI	FALSE	5	TRUE FALSE	0	0	0

Largest mismatch:

[1] 0.02500693409

7.2 Mismatch confidence intervals / boundaries

7.2.1 2 stages

When concluding for futility:

	hypo	missing	ar	binding	fixC	method 1	method 2	method 3
1:	power	TRUE	10	TRUE	FALSE	0	0	0
2:	typeI	TRUE	10	TRUE	FALSE	0	0	0
3:	power	TRUE	5	TRUE	FALSE	0	0	0
4:	typeI	TRUE	5	TRUE	FALSE	0	0	0
5:	power	TRUE	10	TRUE	TRUE	0	0	0
6:	typeI	TRUE	10	TRUE	TRUE	0	0	0
7:	power	TRUE	5	TRUE	TRUE	0	0	0
8:	typeI	TRUE	5	TRUE	TRUE	0	0	0
9:	power	TRUE	10	FALSE	TRUE	O (NA: 37.73%)	0 (NA: 34.80%)	0 (NA: 35.93%)
10:	typeI	TRUE	10	FALSE	TRUE	0 (NA: 0.24%)	0 (NA: 0.24%)	0 (NA: 0.37%)
11:	power	TRUE	5	FALSE	TRUE	0 (NA: 35.87%)	0 (NA: 33.40%)	0 (NA: 34.53%)
12:	typeI	TRUE	5	FALSE	TRUE	O (NA: 0.09%)	0 (NA: 0.09%)	0 (NA: 0.13%)
13:	power	TRUE	10	FALSE	FALSE	O (NA: 34.13%)	0 (NA: 35.61%)	0 (NA: 35.93%)
14:	typeI	TRUE	10	FALSE	FALSE	0 (NA: 0.08%)	0 (NA: 0.10%)	0 (NA: 0.37%)
15:	power	TRUE	5	FALSE	FALSE	O (NA: 33.44%)	0 (NA: 33.62%)	0 (NA: 34.53%)
16:	typeI	TRUE	5	FALSE	FALSE	0 (NA: 0.02%)	0 (NA: 0.02%)	0 (NA: 0.13%)
17:	power	FALSE	5	TRUE	FALSE	0	0	0
18:	typeI	FALSE	5	TRUE	FALSE	0	0	0

When concluding for efficacy:

	hypo	missing	ar	${\tt binding}$	fixC	${\tt method}$	1	${\tt method}$	2	${\tt method}$	3
1:	power	TRUE	10	TRUE	FALSE		0		0		0
2:	typeI	TRUE	10	TRUE	FALSE		0		0		0
3:	power	TRUE	5	TRUE	FALSE		0		0		0
4:	typeI	TRUE	5	TRUE	FALSE		0		0		0
5:	power	TRUE	10	TRUE	TRUE		0		0		0
6:	typeI	TRUE	10	TRUE	TRUE		0		0		0
7:	power	TRUE	5	TRUE	TRUE		0		0		0
8:	typeI	TRUE	5	TRUE	TRUE		0		0		0
9:	power	TRUE	10	FALSE	TRUE		0		0		0
10:	typeI	TRUE	10	FALSE	TRUE		0		0		0
11:	power	TRUE	5	FALSE	TRUE		0		0		0
12:	typeI	TRUE	5	FALSE	TRUE		0		0		0
13:	power	TRUE	10	FALSE	FALSE		0		0		0
14:	typeI	TRUE	10	FALSE	FALSE		0		0		0
15:	power	TRUE	5	FALSE	FALSE		0		0		0
16:	typeI	TRUE	5	FALSE	FALSE		0		0		0

17: power	FALSE	5	TRUE FALSE	0	0	0
18: typeI	FALSE	5	TRUE FALSE	0	0	0

7.2.2 3 stages

When concluding for futility:

	hypo	missing	ar	binding	fixC	method 1	method 2	method 3
1:	power	TRUE	10	TRUE	FALSE	0	0	0
2:	typeI	TRUE	10	TRUE	FALSE	0	0	0
3:	power	TRUE	5	TRUE	FALSE	0	0	0
4:	typeI	TRUE	5	TRUE	FALSE	0	0	0
5:	power	TRUE	10	TRUE	TRUE	0	0	0
6:	typeI	TRUE	10	TRUE	TRUE	0	0	0
7:	power	TRUE	5	TRUE	TRUE	0	0	0
8:	typeI	TRUE	5	TRUE	TRUE	0	0	0
9:	power	TRUE	10	FALSE	TRUE	0 (NA: 48.25%)	0 (NA: 45.63%)	O (NA: 47.50%)
10:	typeI	TRUE	10	FALSE	TRUE	0 (NA: 0.34%)	0 (NA: 0.31%)	O (NA: 0.55%)
11:	power	TRUE	5	FALSE	TRUE	0 (NA: 46.92%)	0 (NA: 43.72%)	0 (NA: 45.38%)
12:	typeI	TRUE	5	FALSE	TRUE	O (NA: 0.09%)	0 (NA: 0.08%)	O (NA: 0.15%)
13:	power	TRUE	10	FALSE	FALSE	0 (NA: 45.00%)	0 (NA: 47.11%)	O (NA: 47.50%)
14:	typeI	TRUE	10	FALSE	FALSE	0 (NA: 0.19%)	0 (NA: 0.22%)	O (NA: 0.55%)
15:	power	TRUE	5	FALSE	FALSE	O (NA: 44.89%)	0 (NA: 45.10%)	0 (NA: 45.38%)
16:	typeI	TRUE	5	FALSE	FALSE	O (NA: 0.07%)	O (NA: 0.07%)	0 (NA: 0.15%)
17:	power	FALSE	5	TRUE	FALSE	0	0	0
18:	typeI	FALSE	5	TRUE	FALSE	0	0	0

When concluding for efficacy:

	hypo	missing	ar	binding	fixC	${\tt method}$	1	${\tt method}$	2	${\tt method}$	3
1:	power	TRUE	10	TRUE	FALSE		0		0		0
2:	typeI	TRUE	10	TRUE	FALSE		0		0		0
3:	power	TRUE	5	TRUE	FALSE		0		0		0
4:	typeI	TRUE	5	TRUE	FALSE		0		0		0
5:	power	TRUE	10	TRUE	TRUE		0		0		0
6:	typeI	TRUE	10	TRUE	TRUE		0		0		0
7:	power	TRUE	5	TRUE	TRUE		0		0		0
8:	typeI	TRUE	5	TRUE	TRUE		0		0		0
9:	power	TRUE	10	FALSE	TRUE		0		0		0
10:	typeI	TRUE	10	FALSE	TRUE		0		0		0
11:	power	TRUE	5	FALSE	TRUE		0		0		0
12:	typeI	TRUE	5	FALSE	TRUE		0		0		0
13:	power	TRUE	10	FALSE	FALSE		0		0	0.01	L%
14:	typeI	TRUE	10	FALSE	FALSE		0		0		0

15:	power	TRUE	5	FALSE	FALSE	0	0	0
16:	typeI	TRUE	5	FALSE	FALSE	0	0	0
17:	power	FALSE	5	TRUE	FALSE	0	0	0
18:	typeI	FALSE	5	TRUE	FALSE	0	0	0

[1] -1.665334537e-16

7.3 Range of p-values

7.3.1 2 stages

	missing	binding	fixC	ar	hypo	method 1	method 2	method 3
1:	TRUE	TRUE	FALSE	10	power	[0;0.9163]	[0;0.9163]	[0;0.9163]
2:	TRUE	TRUE	FALSE	10	typeI	[0;1]	[0;1]	[0;1]
3:	TRUE	TRUE	FALSE	5	power	[0;0.9194]	[0;0.9194]	[0;0.9202]
4:	TRUE	TRUE	FALSE	5	typeI	[0;1]	[0;1]	[0;1]
5:	TRUE	TRUE	TRUE	10	power	[0;0.9534]	[0;0.9557]	[0;0.9163]
6:	TRUE	TRUE	TRUE	10	typeI	[0;1]	[0;1]	[0;1]
7:	TRUE	TRUE	TRUE	5	power	[0;0.9606]	[0;0.9624]	[0;0.9202]
8:	TRUE	TRUE	TRUE	5	typeI	[1e-04;1]	[1e-04;1]	[0;1]
9:	TRUE	FALSE	TRUE	10	power	[0;1]	[0;1]	[0;1]
10:	TRUE	FALSE	TRUE	10	typeI	[3e-04;1]	[4e-04;1]	[2e-04;1]
11:	TRUE	FALSE	TRUE	5	power	[0;1]	[0;1]	[0;1]
12:	TRUE	FALSE	TRUE	5	typeI	[3e-04;1]	[3e-04;1]	[1e-04;1]
13:	TRUE	FALSE	FALSE	10	power	[0;1]	[0;1]	[0;1]
14:	TRUE	FALSE	FALSE	10	typeI	[1e-04;1]	[1e-04;1]	[2e-04;1]
15:	TRUE	FALSE	FALSE	5	power	[0;1]	[0;1]	[0;1]
16:	TRUE	FALSE	FALSE	5	typeI	[1e-04;1]	[1e-04;1]	[1e-04;1]
17:	FALSE	TRUE	FALSE	5	power	[0;0.867]	[0;0.867]	[0;0.882]
18:	FALSE	TRUE	FALSE	5	typeI	[0;0.9999]	[0;0.9999]	[0;1]

7.3.2 3 stages

	missing	binding	fixC	ar	hypo	method 1	method 2	method 3
1:	TRUE	TRUE	FALSE	10	power	[0;0.8665]	[0;0.8663]	[0;0.9035]
2:	TRUE	TRUE	FALSE	10	typeI	[1e-04;0.9999]	[1e-04;0.9999]	[2e-04;1]
3:	TRUE	TRUE	FALSE	5	power	[0;0.8959]	[0;0.8959]	[0;0.9405]
4:	TRUE	TRUE	FALSE	5	typeI	[0;0.9998]	[0;0.9998]	[0;0.9999]
5:	TRUE	TRUE	TRUE	10	power	[0;0.935]	[0;0.9394]	[0;0.9035]
6:	TRUE	TRUE	TRUE	10	typeI	[3e-04;1]	[3e-04;1]	[2e-04;1]
7:	TRUE	TRUE	TRUE	5	power	[0;0.965]	[0;0.9659]	[0;0.9405]
8:	TRUE	TRUE	TRUE	5	typeI	[0;1]	[1e-04;1]	[0;0.9999]
9:	TRUE	FALSE	TRUE	10	power	[0;1]	[0;1]	[0;1]
10:	TRUE	FALSE	TRUE	10	typeI	[4e-04;1]	[5e-04;1]	[3e-04;1]
11:	TRUE	FALSE	TRUE	5	power	[0;1]	[0;1]	[0;1]
12:	TRUE	FALSE	TRUE	5	typeI	[3e-04;1]	[4e-04;1]	[1e-04;1]
13:	TRUE	FALSE	FALSE	10	power	[0;1]	[0;1]	[0;1]
14:	TRUE	FALSE	FALSE	10	typeI	[1e-04;1]	[1e-04;1]	[3e-04;1]
15:	TRUE	FALSE	FALSE	5	power	[0;1]	[0;1]	[0;1]
16:	TRUE	FALSE	FALSE	5	typeI	[1e-04;1]	[1e-04;1]	[1e-04;1]
17:	FALSE	TRUE	FALSE	5	power	[0;0.8745]	[0;0.8745]	[0;0.906]
18:	FALSE	TRUE	FALSE	5	typeI	[0;0.9998]	[0;0.9998]	[0;0.9999]

8 Coverage

8.1 2 stages

	hypo	missing	ar	binding	fixC		me	ethod 1	1 method			2 method 3		
1:	power	FALSE	5	TRUE	FALSE			95.32%			95.35%			95.50%
2:	power	TRUE	5	FALSE	FALSE	96.06%	(NA:	3.12%)	96.06%	(NA:	3.14%)	96.26%	(NA:	2.86%)
3:	power	TRUE	5	FALSE	TRUE	97.57%	(NA:	3.47%)	97.56%	(NA:	3.18%)	96.26%	(NA:	2.86%)
4:	power	TRUE	5	TRUE	FALSE			94.68%			94.69%			94.86%
5:	power	TRUE	5	TRUE	TRUE			95.98%			95.90%			94.86%
6:	power	TRUE	10	FALSE	FALSE	96.16%	(NA:	3.16%)	96.22%	(NA:	3.32%)	95.96%	(NA:	2.52%)
7:	power	TRUE	10	FALSE	TRUE	97.11%	(NA:	3.70%)	97.07%	(NA:	3.36%)	95.96%	(NA:	2.52%)
8:	power	TRUE	10	TRUE	FALSE			94.76%			94.75%			95.04%
9:	power	TRUE	10	TRUE	TRUE			95.55%			95.67%			95.04%
10:	typeI	FALSE	5	TRUE	FALSE			94.96%			94.96%			94.30%
11:	typeI	TRUE	5	FALSE	FALSE	95.02%	(NA:	0.02%)	95.02%	(NA:	0.02%)	95.11%	(NA:	0.12%)
12:	typeI	TRUE	5	FALSE	TRUE	95.09%	(NA:	0.09%)	95.08%	(NA:	0.09%)	95.11%	(NA:	0.12%)
13:	typeI	TRUE	5	TRUE	FALSE			94.63%			94.63%			94.19%
14:	typeI	TRUE	5	TRUE	TRUE			91.68%			91.52%			94.19%
15:	typeI	TRUE	10	FALSE	FALSE	95.09%	(NA:	0.08%)	95.11%	(NA:	0.10%)	95.29%	(NA:	0.35%)
16:	typeI	TRUE	10	FALSE	TRUE	95.23%	(NA:	0.23%)	95.22%	(NA:	0.23%)	95.29%	(NA:	0.35%)
17:	typeI	TRUE	10	TRUE	FALSE			94.56%			94.57%			94.17%
18:	typeI	TRUE	10	TRUE	TRUE			92.67%			92.39%			94.17%

Average width of the confidence intervals

	hypo	${\tt missing}$	ar	${\tt binding}$	fixC	method 1	method 2	method 3
1:	power	FALSE	5	TRUE	FALSE	1.0532	1.0533	1.053
2:	power	TRUE	5	FALSE	FALSE	1.0453	1.0453	1.045
3:	power	TRUE	5	FALSE	TRUE	1.0520	1.0526	1.045
4:	power	TRUE	5	TRUE	FALSE	1.0520	1.0520	1.051
5:	power	TRUE	5	TRUE	TRUE	1.0586	1.0587	1.051
6:	power	TRUE	10	FALSE	FALSE	1.0559	1.0558	1.051
7:	power	TRUE	10	FALSE	TRUE	1.0638	1.0649	1.051
8:	power	TRUE	10	TRUE	FALSE	1.0627	1.0631	1.058
9:	power	TRUE	10	TRUE	TRUE	1.0708	1.0712	1.058
10:	typeI	FALSE	5	TRUE	FALSE	1.0444	1.0443	1.047
11:	typeI	TRUE	5	FALSE	FALSE	0.9995	0.9995	1.014
12:	typeI	TRUE	5	FALSE	TRUE	0.9994	0.9996	1.014
13:	typeI	TRUE	5	TRUE	FALSE	1.0431	1.0429	1.047
14:	typeI	TRUE	5	TRUE	TRUE	1.0396	1.0406	1.047
15:	typeI	TRUE	10	FALSE	FALSE	1.0009	1.0009	1.044
16:	typeI	TRUE	10	FALSE	TRUE	1.0005	1.0005	1.044
17:	typeI	TRUE	10	TRUE	FALSE	1.0469	1.0457	1.057
18:	typeI	TRUE	10	TRUE	TRUE	1.0412	1.0425	1.057

Average ratio between the length of the MUE CIs vs. the ML CIs

	hypo	missing	ar	binding	fixC	method 1	method 2	method 3
1:	power	FALSE	5	TRUE	FALSE	1.0579	1.0580	1.058
2:	power	TRUE	5	FALSE	FALSE	1.0539	1.0539	1.053
3:	power	TRUE	5	FALSE	TRUE	1.0602	1.0604	1.053
4:	power	TRUE	5	TRUE	FALSE	1.0574	1.0574	1.057
5:	power	TRUE	5	TRUE	TRUE	1.0633	1.0632	1.057
6:	power	TRUE	10	FALSE	FALSE	1.0603	1.0603	1.056
7:	power	TRUE	10	FALSE	TRUE	1.0680	1.0688	1.056
8:	power	TRUE	10	TRUE	FALSE	1.0652	1.0656	1.060
9:	power	TRUE	10	TRUE	TRUE	1.0727	1.0730	1.060
10:	typeI	FALSE	5	TRUE	FALSE	1.0506	1.0506	1.054
11:	typeI	TRUE	5	FALSE	FALSE	0.9995	0.9995	1.014
12:	typeI	TRUE	5	FALSE	TRUE	0.9996	0.9997	1.014
13:	typeI	TRUE	5	TRUE	FALSE	1.0499	1.0497	1.054
14:	typeI	TRUE	5	TRUE	TRUE	1.0467	1.0477	1.054
15:	typeI	TRUE	10	FALSE	FALSE	1.0009	1.0010	1.045
16:	typeI	TRUE	10	FALSE	TRUE	1.0008	1.0009	1.045
17:	typeI	TRUE	10	TRUE	FALSE	1.0502	1.0490	1.061
18:	typeI	TRUE	10	TRUE	TRUE	1.0448	1.0461	1.061

8.2 3 stages

	hypo	missing	ar	binding	fixC	method 1			me	ethod 2		method 3		
1:	power	FALSE	5	TRUE	FALSE			94.74%	94.73%	(NA:	0.02%)	95.00%	(NA:	0.01%)
2:	power	TRUE	5	FALSE	FALSE	95.81%	(NA:	4.35%)	95.85%	(NA:	4.37%)	95.85%	(NA:	3.85%)
3:	power	TRUE	5	FALSE	TRUE	97.75%	(NA:	4.72%)	97.73%	(NA:	4.32%)	95.86%	(NA:	3.85%)
4:	power	TRUE	5	TRUE	FALSE	95.11%	(NA:	0.04%)	95.14%	(NA:	0.04%)	95.32%	(NA:	0.03%)
5:	power	TRUE	5	TRUE	TRUE	96.03%	(NA:	0.03%)	96.15%	(NA:	0.03%)	95.32%	(NA:	0.03%)
6:	power	TRUE	10	FALSE	FALSE	95.81%	(NA:	4.23%)	95.89%	(NA:	4.48%)	95.54%	(NA:	3.21%)
7:	power	TRUE	10	FALSE	TRUE	97.48%	(NA:	4.82%)	97.50%	(NA:	4.44%)	95.52%	(NA:	3.21%)
8:	power	TRUE	10	TRUE	FALSE			95.15%			95.16%			95.33%
9:	power	TRUE	10	TRUE	TRUE			95.97%			95.91%			95.33%
10:	typeI	FALSE	5	TRUE	FALSE	94.80%	(NA:	0.13%)	94.80%	(NA:	0.13%)	94.19%	(NA:	0.15%)
11:	typeI	TRUE	5	FALSE	FALSE	94.78%	(NA:	0.30%)	94.83%	(NA:	0.35%)	94.71%	(NA:	0.40%)
12:	typeI	TRUE	5	FALSE	TRUE	94.80%	(NA:	0.32%)	94.79%	(NA:	0.30%)	94.75%	(NA:	0.40%)
13:	typeI	TRUE	5	TRUE	FALSE	95.05%	(NA:	0.12%)	95.05%	(NA:	0.12%)	94.20%	(NA:	0.16%)
14:	typeI	TRUE	5	TRUE	TRUE	89.80%	(NA:	0.13%)	89.53%	(NA:	0.15%)	94.20%	(NA:	0.15%)
15:	typeI	TRUE	10	FALSE	FALSE	94.85%	(NA:	0.42%)	94.90%	(NA:	0.47%)	95.13%	(NA:	0.77%)
16:	typeI	TRUE	10	FALSE	TRUE	95.00%	(NA:	0.57%)	95.01%	(NA:	0.59%)	95.12%	(NA:	0.79%)
17:	typeI	TRUE	10	TRUE	FALSE	95.02%	(NA:	0.22%)	94.97%	(NA:	0.14%)	94.30%	(NA:	0.09%)
18:	typeI	TRUE	10	TRUE	TRUE	91.52%	(NA:	0.17%)	91.12%	(NA:	0.14%)	94.30%	(NA:	0.11%)

Average width of the confidence intervals

```
hypo missing ar binding fixC method 1 method 2 method 3
 1: power
            FALSE
                   5
                         TRUE FALSE
                                       1.0789
                                                1.0790
                                                           1.077
 2: power
             TRUE
                   5
                                                1.0707
                                                           1.069
                        FALSE FALSE
                                       1.0707
3: power
             TRUE
                   5
                        FALSE
                              TRUE
                                       1.0757
                                                1.0757
                                                           1.069
4: power
             TRUE
                   5
                         TRUE FALSE
                                       1.0782
                                                1.0783
                                                           1.077
5: power
             TRUE
                   5
                         TRUE
                              TRUE
                                       1.0834
                                                1.0830
                                                           1.077
6: power
             TRUE 10
                        FALSE FALSE
                                       1.1011
                                                1.1012
                                                           1.096
7: power
             TRUE 10
                        FALSE
                                                           1.096
                              TRUE
                                       1.1101
                                                1.1110
8: power
             TRUE 10
                         TRUE FALSE
                                       1.1085
                                                1.1087
                                                           1.104
9: power
             TRUE 10
                         TRUE
                              TRUE
                                       1.1176
                                                1.1176
                                                           1.104
10: typeI
            FALSE
                         TRUE FALSE
                                                1.0788
                                                           1.084
                   5
                                       1.0788
11: typeI
             TRUE
                   5
                        FALSE FALSE
                                       0.9997
                                                0.9997
                                                           1.018
12: typeI
             TRUE
                   5
                        FALSE
                              TRUE
                                       0.9998
                                                0.9998
                                                           1.018
13: typeI
             TRUE
                   5
                         TRUE FALSE
                                       1.0743
                                                1.0742
                                                           1.079
14: typeI
             TRUE
                   5
                         TRUE
                              TRUE
                                       1.0755
                                                1.0764
                                                           1.079
15: typeI
             TRUE 10
                        FALSE FALSE
                                       0.9995
                                                0.9995
                                                           1.061
16: typeI
             TRUE 10
                        FALSE
                              TRUE
                                       0.9990
                                                0.9994
                                                           1.061
17: typeI
             TRUE 10
                         TRUE FALSE
                                       1.1013
                                                1.1005
                                                           1.114
18: typeI
             TRUE 10
                         TRUE
                               TRUE
                                       1.0942
                                                1.0953
                                                           1.114
```

Average ratio between the length of the MUE CIs vs. the ML CIs

	hypo	missing	ar	binding	fixC	method 1	${\tt method}\ 2$	method 3
1:	power	FALSE	5	TRUE	FALSE	1.0825	1.0826	1.081
2:	power	TRUE	5	FALSE	FALSE	1.0776	1.0776	1.076
3:	power	TRUE	5	FALSE	TRUE	1.0811	1.0808	1.076
4:	power	TRUE	5	TRUE	FALSE	1.0827	1.0828	1.081
5:	power	TRUE	5	TRUE	TRUE	1.0859	1.0854	1.081
6:	power	TRUE	10	FALSE	FALSE	1.1040	1.1041	1.098
7:	power	TRUE	10	FALSE	TRUE	1.1117	1.1120	1.098
8:	power	TRUE	10	TRUE	FALSE	1.1100	1.1103	1.105
9:	power	TRUE	10	TRUE	TRUE	1.1173	1.1171	1.105
10:	typeI	FALSE	5	TRUE	FALSE	1.0854	1.0854	1.091
11:	typeI	TRUE	5	FALSE	FALSE	0.9996	0.9996	1.018
12:	typeI	TRUE	5	FALSE	TRUE	0.9998	0.9997	1.018
13:	typeI	TRUE	5	TRUE	FALSE	1.0815	1.0814	1.087
14:	typeI	TRUE	5	TRUE	TRUE	1.0844	1.0851	1.087
15:	typeI	TRUE	10	FALSE	FALSE	0.9995	0.9995	1.062
16:	typeI	TRUE	10	FALSE	TRUE	0.9994	0.9996	1.062
17:	typeI	TRUE	10	TRUE	FALSE	1.1060	1.1052	1.120
18:	tvpeI	TRUE	10	TRUE	TRUE	1.1006	1.1013	1.120

9 Percentage of missing values (2 stages)

At the first interim

- pc.all percentage of observations with full data (with respect to all observations, i.e. patients with baseline measurement)
- pc.missing3 percentage of observations missing the final outcome but with intermediate outcome value and baseline.
- pc.missing23 percentage of observations with only baseline value

Here only for method 1 - values are very similar between different methods:

	${\tt method}$	missing	ar	hypo	fixC	${\tt binding}$	N	<pre>pc.all</pre>	${\tt pc.missing3}$	pc.missing23
1:	1	TRUE	5	power	FALSE	TRUE	10000	78.81	9.833	11.360
2:	1	TRUE	5	typeI	FALSE	TRUE	10000	78.81	9.833	11.360
3:	1	TRUE	5	power	TRUE	TRUE	10000	78.81	9.833	11.360
4:	1	TRUE	5	typeI	TRUE	TRUE	10000	78.81	9.833	11.360
5:	1	TRUE	5	power	TRUE	FALSE	9900	78.73	9.861	11.410
6:	1	TRUE	5	typeI	TRUE	FALSE	10000	78.73	9.860	11.410
7:	1	TRUE	5	power	FALSE	FALSE	9900	78.73	9.861	11.410
8:	1	TRUE	5	typeI	FALSE	FALSE	10000	78.73	9.860	11.410
9:	1	FALSE	5	power	FALSE	TRUE	10000	86.82	6.570	6.610
10:	1	FALSE	5	typeI	FALSE	TRUE	9900	86.82	6.570	6.609
11:	1	TRUE	10	power	FALSE	TRUE	10000	70.41	13.802	15.791
12:	1	TRUE	10	typeI	FALSE	TRUE	10000	70.41	13.802	15.791
13:	1	TRUE	10	power	TRUE	TRUE	10000	70.41	13.802	15.791
14:	1	TRUE	10	typeI	TRUE	TRUE	10000	70.41	13.802	15.791
15:	1	TRUE	10	power	TRUE	FALSE	9900	70.42	13.807	15.776
16:	1	TRUE	10	typeI	TRUE	FALSE	10000	70.42	13.806	15.776
17:	1	TRUE	10	power	FALSE	FALSE	9900	70.42	13.807	15.776
18:	1	TRUE	10	typeI	FALSE	FALSE	10000	70.42	13.806	15.776

10 Information

10.1 2 stages

Percentage of information for method 1^5 :

```
scenario missing binding fixC ar interim decision final
                    TRUE FALSE 10
                                     54.92
                                              77.30 103.07
       1
            TRUE
       2
            TRUE
                    TRUE FALSE 10
                                     54.92
                                              77.05 102.89
       3
                    TRUE FALSE 5
                                              64.89 103.08
            TRUE
                                     53.43
       4
            TRUE
                    TRUE FALSE 5
                                     53.43
                                              64.61 102.94
       5
            TRUE
                    TRUE
                          TRUE 10
                                     54.92
                                              77.30 103.07
       6
            TRUE
                    TRUE
                          TRUE 10
                                     54.92
                                              77.05 102.89
       7
                    TRUE
                                              64.89 103.08
            TRUE
                          TRUE 5
                                     53.43
       8
                    TRUE
                          TRUE 5
                                              64.61 102.94
            TRUE
                                     53.43
       9
            TRUE
                   FALSE
                          TRUE 10
                                     54.72
                                              76.74 102.63
                   FALSE
                          TRUE 10
                                              76.11 103.18
      10
            TRUE
                                     54.71
            TRUE
                   FALSE
                          TRUE
                                     53.25
                                              64.47 102.68
      11
      12
            TRUE
                   FALSE
                          TRUE 5
                                     53.25
                                              64.35 103.18
      13
            TRUE
                   FALSE FALSE 10
                                     54.72
                                              76.74 102.63
                                     54.71
                                              76.11 103.18
      14
            TRUE
                   FALSE FALSE 10
                   FALSE FALSE 5
                                              64.47 102.68
      15
            TRUE
                                     53.25
            TRUE
                   FALSE FALSE 5
                                     53.25
                                              64.35 103.18
      16
                    TRUE FALSE 5
                                              64.38 100.18
      17
           FALSE
                                     52.13
      18
           FALSE
                    TRUE FALSE 5
                                     52.12
                                              64.19 99.81
```

Similar results for other methods.

10.2 3 stages

Percentage of information for method 1^6 :

scenario	missing	binding	fixC	ar	interim1	decision1	interim2	decision2	final3
1	TRUE	TRUE	FALSE	10	39.08	61.13	64.87	86.70	102.65
2	TRUE	TRUE	FALSE	10	39.08	61.04	64.66	86.27	102.16
3	TRUE	TRUE	FALSE	5	37.69	48.99	63.40	74.47	102.73
4	TRUE	TRUE	FALSE	5	37.69	48.93	63.17	74.09	102.19
5	TRUE	TRUE	TRUE	10	39.08	61.13	64.87	86.70	102.65
6	TRUE	TRUE	TRUE	10	39.08	61.04	64.66	86.27	102.16
7	TRUE	TRUE	TRUE	5	37.69	48.99	63.40	74.47	102.73
8	TRUE	TRUE	TRUE	5	37.69	48.93	63.17	74.09	102.19
9	TRUE	FALSE	TRUE	10	38.85	60.43	64.52	85.88	102.34
10	TRUE	FALSE	TRUE	10	38.85	61.18	64.76	90.06	102.97

⁵average over the reached stages

 $^{^6}$ average over the reached stages

11	TRUE	FALSE	TRUE	5	37.47	48.55	63.08	73.93 102.43
12	TRUE	FALSE	TRUE	5	37.47	49.96	63.30	75.12 103.00
13	TRUE	FALSE	FALSE	10	38.85	60.43	64.52	85.88 102.34
14	TRUE	FALSE	FALSE	10	38.85	61.18	64.76	90.06 102.97
15	TRUE	FALSE	FALSE	5	37.47	48.55	63.08	73.93 102.43
16	TRUE	FALSE	FALSE	5	37.47	49.96	63.30	75.12 103.00
17	FALSE	TRUE	FALSE	5	36.89	49.06	61.81	73.70 99.93
18	FALSE	TRUE	FALSE	5	36.89	48.91	61.69	73.42 99.67