

(\* Mathematica Code for determination of optimal decision thresholds

[aC1,bC1]:	prior interval for event rates in G_1 in the control group
[aC2,bC2]:	prior interval for event rates in G_2 in the control group
[aT1,bT1]:	prior interval for event rates in G_1 in the treatment group
[aT2,bT2]:	prior interval for event rates in G_2 in the treatment group
n:	number of patients per group and stage
prev:	prevalence of G_1
tau0:	relevance threshold for G_0
tau1:	relevance threshold for G_1
c0:	decision threshold for G_0 to be optimized
c1:	decision threshold for G_1 to be optimized
g0[]:	derivative of the expected loss given G_0 is wrongly selected
h0[]:	derivative of the expected loss given G_0 is wrongly not selected
g1[]:	derivative of the expected loss given G_1 is wrongly selected
h1[]:	derivative of the expected loss given G_1 is wrongly not selected
f0[]:	derivative of the expected loss given a false decision concerning G_0 occurred
f1[]:	derivative of the expected loss given a false decision concerning G_1 occurred
b[]:	function that rounds the results on four digits and maps values smaller than -1 to -1 and values larger than 1 to 1, respectively
opt0[]:	yields the root of f0, i.e. the optimal threshold c0*
opt1[]:	yields the root of f1, i.e. the optimal threshold c1*
opt[]:	yields [c0*,c1*]

Local adaptive method was applied for numerical integration with precision of 8 digits.

Secant method was applied for numerical root solving with precision of 3 digits.\*)

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g0[aC1_?NumericQ, bC1_?NumericQ, aC2_?NumericQ, bC2_?NumericQ,
  aT1_?NumericQ, bT1_?NumericQ, aT2_?NumericQ, bT2_?NumericQ,
  tau0_?NumericQ, n_?NumericQ, prev_?NumericQ, c0_?NumericQ] :=
  NIntegrate[-((prev * (pT1 - pC1) + (1 - prev) * (pT2 - pC2) - tau0))^(2) *
    Boole[((prev * (pT1 - pC1) + (1 - prev) * (pT2 - pC2))) < tau0] * Exp[-(c0 -
      (prev * (pT1 - pC1) + (1 - prev) * (pT2 - pC2)))^2 /
    (2 * ((prev * (pT1 (1 - pT1) + pC1 (1 - pC1)) +
      (1 - prev) * (pT2 (1 - pT2) + pC2 (1 - pC2))) / n))],
  {pT1, aT1, bT1}, {pC1, aC1, bC1}, {pT2, aT2, bT2}, {pC2, aC2, bC2},
  Method → "LocalAdaptive",
  PrecisionGoal → 8]

h0[aC1_?NumericQ, bC1_?NumericQ, aC2_?NumericQ, bC2_?NumericQ,
  aT1_?NumericQ, bT1_?NumericQ, aT2_?NumericQ, bT2_?NumericQ,
  tau0_?NumericQ, n_?NumericQ, prev_?NumericQ, c0_?NumericQ] :=
  NIntegrate[+((prev * (pT1 - pC1) + (1 - prev) * (pT2 - pC2) - tau0))^(2) *
    Boole[((prev * (pT1 - pC1) + (1 - prev) * (pT2 - pC2))) > tau0] * Exp[-(c0 -
      (prev * (pT1 - pC1) + (1 - prev) * (pT2 - pC2)))^2 /
    (2 * ((prev * (pT1 (1 - pT1) + pC1 (1 - pC1)) +
      (1 - prev) * (pT2 (1 - pT2) + pC2 (1 - pC2))) / n))],
  {pT1, aT1, bT1}, {pC1, aC1, bC1}, {pT2, aT2, bT2}, {pC2, aC2, bC2},
  Method → "LocalAdaptive",
  PrecisionGoal → 8]

g1[aC1_?NumericQ, bC1_?NumericQ, aT1_?NumericQ, bT1_?NumericQ,
  tau1_?NumericQ, n_?NumericQ, prev_?NumericQ, c1_?NumericQ] :=
  NIntegrate[-((pT1 - pC1) - tau1)^2 * Boole[(pT1 - pC1) < tau1] * Exp[-(c1 -
    (pT1 - pC1))^2 / (2 * ((pT1 (1 - pT1) + pC1 (1 - pC1)) / (prev * n))],
  {pT1, aT1, bT1}, {pC1, aC1, bC1}, Method → "LocalAdaptive", PrecisionGoal → 8]

h1[aC1_?NumericQ, bC1_?NumericQ, aT1_?NumericQ, bT1_?NumericQ,
  tau1_?NumericQ, n_?NumericQ, prev_?NumericQ, c1_?NumericQ] :=
  NIntegrate[+((pT1 - pC1) - tau1)^2 * Boole[(pT1 - pC1) > tau1] * Exp[-(c1 -
    (pT1 - pC1))^2 / (2 * ((pT1 (1 - pT1) + pC1 (1 - pC1)) / (prev * n))],
  {pT1, aT1, bT1}, {pC1, aC1, bC1}, Method → "LocalAdaptive", PrecisionGoal → 8]

f0[aC1_, bC1_, aC2_, bC2_, aT1_, bT1_, aT2_, bT2_, tau0_, n_, prev_, c0_] :=
  g0[aC1, bC1, aC2, bC2, aT1, bT1, aT2, bT2, tau0, n, prev, c0] +
  h0[aC1, bC1, aC2, bC2, aT1, bT1, aT2, bT2, tau0, n, prev, c0]

f1[aC1_, bC1_, aT1_, bT1_, tau1_, n_, prev_, c1_] :=
  g1[aC1, bC1, aT1, bT1, tau1, n, prev, c1] + h1[aC1, bC1, aT1, bT1, tau1, n, prev, c1]

b[x_] := Round[Sign[x] * Min[Abs[x], 1], 0.0001]

opt0[aC1_?NumericQ, bC1_?NumericQ, aC2_?NumericQ,
  bC2_?NumericQ, aT1_?NumericQ, bT1_?NumericQ, aT2_?NumericQ,
  bT2_?NumericQ, tau0_?NumericQ, n_?NumericQ, prev_?NumericQ] :=
  b[c0 /. FindRoot[f0[aC1, bC1, aC2, bC2, aT1, bT1, aT2, bT2, tau0, n, prev, c0],
    {c0, -1, 1, -2, 2}], PrecisionGoal → 3]]

opt1[aC1_?NumericQ, bC1_?NumericQ, aT1_?NumericQ,
  bT1_?NumericQ, tau1_?NumericQ, n_?NumericQ, prev_?NumericQ] :=
  b[c1 /. FindRoot[f1[aC1, bC1, aT1, bT1, tau1, n, prev, c1],
    {c1, -1, 1, -2, 2}], PrecisionGoal → 3]]

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opt[pC11_, pC1h_, pC21_, pC2h_, pT11_, pT1h_, pT21_, pT2h_, tau0_, tau1_, n_,
 prev_] := {opt0[pC11, pC1h, pC21, pC2h, pT11, pT1h, pT21, pT2h, tau0, n, prev], 
 opt1[pC11, pC1h, pT11, pT1h, tau1, n, prev]}

(* some example *)

opt[0.05, 0.35, 0.2, 0.5, 0.3, 0.6, 0.2, 0.5, 0.05, 0.1, 100, 0.25]
{0.0455, -0.0826}
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