Admissible Multinomial Trial - IUT design Example

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Multinomial Trial Design with Different Output Options

In the following example, we provide four design methods for multinomial trial: Minimax (minimize the maximum sample size), Optimal (minimize the expected sample size), Admissible (minimize the Bayesian risk) and Maxpower (maximize the exact power level).

```
source(file="powfun_IUT.r")
source(file="searchfun_IUT_adm.r")
library(clinfun)
```

Single-stage

It should be noted that single-stage design only allows Minimax and Maxpower output methods.

```
# Minimax
IUT.design(method = "s1", s2.rej = 18, t2.rej = 12, n = 80, s2.rej.delta = 1,
   t2.rej.delta = 1, n.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3, p1.t = 0.1,
   output = "minimax")
    p0.s p0.t p1.s p1.t s.rej t.rej N Error Power
## 2 0.15 0.25 0.3 0.1
                           18
                                 11 79 0.0430 0.857
## 3 0.15 0.25 0.3 0.1
                           19
                                 11 79 0.0228 0.825
## 5 0.15 0.25 0.3 0.1
                           18
                                 12 79 0.0430 0.896
## 6 0.15 0.25 0.3 0.1
                           19
                                 12 79 0.0254 0.862
## 8 0.15 0.25 0.3 0.1
                           18
                                 13 79 0.0477 0.919
## 9 0.15 0.25 0.3 0.1
                           19
                                 13 79 0.0477 0.882
##
     user system elapsed
##
     0.49
             0.02
                     0.50
# Maxpower
IUT.design(method = "s1", s2.rej = 18, t2.rej = 12, n = 80, s2.rej.delta = 1,
   t2.rej.delta = 1, n.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3, p1.t = 0.1,
   output = "maxpower")
##
     p0.s p0.t p1.s p1.t s.rej t.rej N Error Power
                            18
                                  13 80 0.048 0.924
## 17 0.15 0.25 0.3 0.1
##
     user system elapsed
     0.56
             0.00
##
                     0.57
```

Two-stage

```
# Minimax
IUT.design(method = "s2.sf", s1.rej = 10, t1.rej = 3, s1.acc = 8, t1.acc = 5,
   s2.rej = 18, t2.rej = 12, n1 = 41, n2 = 41, s1.rej.delta = 1, t1.rej.delta = 1,
   s2.rej.delta = 1, t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3,
   p1.t = 0.1, output = "minimax")
     p0.s p0.t p1.s p1.t s1.rej t1.rej s1.acc t1.acc s2.rej t2.rej N1 N2
##
## 27 0.15 0.25 0.3 0.1
                                     4
                                            8
                                                   5
                                                                11 41 41
                             11
                                                         19
## 54 0.15 0.25 0.3 0.1
                                            8
                             11
                                     4
                                                   5
                                                         19
                                                                12 41 41
## 81 0.15 0.25 0.3 0.1
                             11
                                     4
                                            8
                                                   5
                                                         19
                                                                13 41 41
      Error Power PET
## 27 0.0476 0.853 0.996 41.2
## 54 0.0476 0.869 0.996 41.2
## 81 0.0476 0.875 0.996 41.2
     user system elapsed
##
    65.09
             0.01
                    65.57
# Optimal
IUT.design(method = "s2.sf", s1.rej = 10, t1.rej = 3, s1.acc = 8, t1.acc = 5,
   s2.rej = 18, t2.rej = 12, n1 = 41, n2 = 41, s1.rej.delta = 1, t1.rej.delta = 1,
   s2.rej.delta = 1, t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3,
 p1.t = 0.1, output = "optimal")
     p0.s p0.t p1.s p1.t s1.rej t1.rej s1.acc t1.acc s2.rej t2.rej N1 N2
## 27 0.15 0.25 0.3 0.1
                             11
                                     4
                                            8
                                                   5
                                                         19
                                                                11 41 41
                                                                12 41 41
## 54 0.15 0.25 0.3 0.1
                                            8
                                                   5
                                                         19
                             11
                                     4
## 81 0.15 0.25 0.3 0.1
                             11
                                            8
                                                   5
                                                         19
                                                                13 41 41
      Error Power PET
## 27 0.0476 0.853 0.996 41.2
## 54 0.0476 0.869 0.996 41.2
## 81 0.0476 0.875 0.996 41.2
##
     user system elapsed
##
    65.13
             0.02
                    65.52
# Admissible
IUT.design(method = "s2.sf", s1.rej = 10, t1.rej = 3, s1.acc = 8, t1.acc = 5,
   s2.rej = 18, t2.rej = 12, n1 = 41, n2 = 41, s1.rej.delta = 1, t1.rej.delta = 1,
   s2.rej.delta = 1, t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3,
   p1.t = 0.1, output = "admissible")
     p0.s p0.t p1.s p1.t s1.rej t1.rej s1.acc t1.acc s2.rej t2.rej N1 N2
## 27 0.15 0.25 0.3 0.1
                             11
                                     4
                                          8
                                                 5 19 11 41 41
      Error Power PET
## 27 0.0476 0.853 0.996 41.2
##
     user system elapsed
    65.45
             0.01
# Maxpower
IUT.design(method = "s2.sf", s1.rej = 10, t1.rej = 3, s1.acc = 8, t1.acc = 5,
   s2.rej = 18, t2.rej = 12, n1 = 41, n2 = 41, s1.rej.delta = 1, t1.rej.delta = 1,
   s2.rej.delta = 1, t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3,
 p1.t = 0.1, output = "maxpower")
```

```
## p0.s p0.t p1.s p1.t s1.rej t1.rej s1.acc t1.acc s2.rej t2.rej N1 N2
## 81 0.15 0.25 0.3 0.1 11 4 8 5 19 13 41 41
## Error Power PET EN
## 81 0.0476 0.875 0.996 41.2
## user system elapsed
## 65.08 0.00 65.30
```

Two-stage early terminate with futility only

```
# Minimax
suppressWarnings(IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17,
   t2.rej = 13, n1 = 41, n2 = 41, s1.acc.delta = 1, t1.acc.delta = 1, s2.rej.delta = 1,
   t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3, p1.t = 0.1, output = "minimax"))
## Error in IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17, : No feasible solution f
## Increase maximum sample size. Current nmax value = 82.
suppressWarnings(IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17,
   t2.rej = 13, n1 = 41, n2 = 41, s1.acc.delta = 1, t1.acc.delta = 1, s2.rej.delta = 1,
   t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3, p1.t = 0.1, output = "optimal"))
## Error in IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17, : No feasible solution f
## Increase maximum sample size. Current nmax value = 82.
# Admissible
suppressWarnings(IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17,
   t2.rej = 13, n1 = 41, n2 = 41, s1.acc.delta = 1, t1.acc.delta = 1, s2.rej.delta = 1,
   t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3, p1.t = 0.1, output = "admissible"))
## Error in IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17, : No feasible solution f
## Increase maximum sample size. Current nmax value = 82.
# Maxpower
suppressWarnings(IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17,
   t2.rej = 13, n1 = 41, n2 = 41, s1.acc.delta = 1, t1.acc.delta = 1, s2.rej.delta = 1,
   t2.rej.delta = 1, p0.s = 0.15, p0.t = 0.25, p1.s = 0.3, p1.t = 0.1, output = "maxpower"))
## Error in IUT.design(method = "s2.f", s1.acc = 7, t1.acc = 5, s2.rej = 17, : No feasible solution f
```

Increase maximum sample size. Current nmax value = 82.