

One-arm binary predictive probability

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Suppose θ is the probability of success in a Bernoulli trial and θ has a $\text{beta}(a, b)$ distribution. The predictive probability of s successes and f failures is

$$\binom{s+f}{s} \frac{B(a+s, b+f)}{B(a, b)}$$

where $B(x, y) = \Gamma(x)\Gamma(y)/\Gamma(x+y)$ is the beta function. For background on the use of predictive probability in clinical trials, see [1].

Let $\varphi(u, v)$ be a stopping rule based on the posterior distribution of θ . The function φ returns 1 if a trial would stop when the posterior distribution of θ has a $\text{beta}(u, v)$ distribution and 0 otherwise. Then the predictive probability of the stopping rule activating after n future observations is

$$\sum_{s=0}^n \binom{s+f}{s} \frac{B(a+s, b+f)}{B(a, b)} \varphi(a+s, b+f)$$

The following R code will compute the predictive probability of an event, such as early stopping, given by a function `fctn` of beta distribution parameters.

```
binarypp <- function(a, b, n, fctn)
{
  sum <- 0.0
  for (s in 0:n)
  {
    f <- n - s
    # work in logs to prevent overflow or underflow
    logprob <- lchoose(s + f, s) + lbeta(a + s, b + f) - lbeta(a, b)
    sum <- sum + exp(logprob)*fctn(a+s, b+f)
  }
  return (sum)
}
```

Here the parameters a and b are the beta parameters for the distribution on θ . This could be the prior distribution, or the posterior distribution after some number of observations. The number of future observations is n .

Suppose a trial would stop if $P(\theta > 0.4) < 0.05$. This could be implemented as follows.

```
stopping.rule <- function(a, b)
{
  # Test whether Prob( X > historical ) < threshold
  historical <- 0.4
  threshold <- 0.05
  return (pbeta(historical, a, b, lower.tail = FALSE) < threshold)
}
```

The predictive probability of a trial stopping would then be found by passing `stopping.rule` to `binarypp`. For example, suppose a trial was designed with a $\text{beta}(0.3, 0.7)$ prior on θ and then observed 30 successes and 50 failures. The predictive probability of stopping the trial after 20 more patients would be

```
binarypp(30.3, 50.7, 20, stopping.rule)
```

1 Reference

[1] John Cook, “Predictive Probability Interim Analysis” available at <http://bit.ly/nxkKg1>.

Expanded address: <https://biostatistics.mdanderson.org/SoftwareDownload/ProductSupportFiles/PredictiveProbabilit/PredictiveInterimAnalysis.pdf>