## ON A SCREENING PROBLEM<sup>1</sup>

## By JOSEPH A. YAHAV

The Hebrew University, Jerusalem

Fisher considered the problem of constructing sequentially a "better" finite population from a given infinite one. The purpose of this paper is to prove the optimality of Fisher's procedure.

1. Introduction. Fisher, in his paper on sequential experimentation [1], considered a problem of sequential screening. In this problem one is interested in selecting a finite number of female mice which have a specific genetic trait. Fisher stated the problem and prescribed a sequential probability ratio test as a solution. It is the purpose of this paper to prove the optimality of Fisher's procedure and to explain and generalize his ideas.

We consider a dichotomized infinite population. The proportion of A-elements is  $\Pi$  and the proportion of  $\overline{A}$ -elements is  $1 - \Pi$ . Given an element, we are not able to tell if it is an A-element or an  $\overline{A}$ -element. However, we can test the element, with a given test procedure, so that we get a positive or a negative result, where

(1.1) 
$$P(+|A) = \alpha, \qquad P(+|\overline{A}) = \beta.$$

We assume  $\Pi$ ,  $\alpha$  and  $\beta$  to be known and  $\beta < \alpha$ . Furthermore, we assume that an element can be tested repeatedly with independent and identically distributed results, conditional on being A or  $\overline{A}$ , satisfying (1.1).

Our target is to construct a finite population, of size N, consisting of elements from the original population so that the proportion of A-elements in the new population exceeds  $\Pi^*$ , where  $\Pi^* > \Pi$ .

As is easily seen, this condition can never be satisfied with a finite number of tests unless  $\alpha = 1$  and  $\beta = 0$ . Hence we reduce our standard somewhat, and ask that, for any element in the newly constructed population, the conditional probability of the element being an A-element (given that the element was selected) is greater or equal to  $\Pi^*$ .

The selection takes place in a sequential manner. There is a fixed cost C (C > 0) for each test; there are no additional costs in the process. The objective is to minimize the expected cost, subject to satisfying the standard.

Fisher prescribed the following procedure: take an element from the original population, keep testing it so long as

(1.2) 
$$\Pi \leq P(A|\text{the results on testing}) < \Pi^*,$$

1140

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and stop the first time the inequality in (1.2) is violated. If  $P(A|\text{the results on testing}) > \Pi^*$ , then select this element. If  $P(A|\text{the results on testing}) < \Pi$ , then reject this element. Continue the testing until N elements are selected.

2. The case N = 1. For N = 1 we have to select one element subject to

(2.1) 
$$P(A|\text{the element was selected}) > \Pi^*$$
.

We restrict ourselves at this stage to procedures that do not permit recall of an element that was tested and rejected. Thus, we test elements  $\varepsilon_1, \varepsilon_2, \cdots$  until an element satisfying (2.1) is selected. A sequential selection procedure is defined by a sequence of bivariate random variables  $(T_1, I_1), (T_2, I_2), \cdots, (T_K, I_K)$ . Where  $T_i$  is a stopping time for tests on element  $\varepsilon_i$  and

(2.2) 
$$I_i = 1$$
 if element  $i$  is selected  $= 0$  if element  $i$  is rejected.

Hence, if  $\varepsilon_K$  is selected, K is a stopping time we have

(2.3) 
$$I_i = 0, \quad i = 1, 2, \dots, K-1; \quad I_K = 1.$$

The total cost is then given by

(2.4) 
$$\operatorname{Cost} = C \cdot \sum_{i=1}^{K} T_{i},$$

where C > 0 is the cost per test.

Our objective is to find stopping times  $T_1, T_2, \cdots$ , and K so that the expected cost is minimized subject to

(2.5) 
$$P(\varepsilon_K \text{ is an } A\text{-element}|I_K = 1) > \Pi^*.$$

Since the problem can be formulated as a negative dynamic programming problem, it is enough to consider stationary procedures. For stationary procedures we have  $(T_i, I_i)$  are i.i.d., so that

(2.6) 
$$E\left[\sum_{i=1}^{K} T_i\right] = E\left[T_1\right] \cdot E\left[K\right].$$

Since  $E[T_1] = \infty$  or  $E[K] = \infty$  implies  $E[\sum_{i=1}^K T_i] = \infty$ , it is enough to exhibit a stationary procedure for which  $E[T_i] < \infty$  and  $E[K] < \infty$  in order to conclude that an optimal stationary procedure (if it exists) satisfies (2.6).

Since the  $I_i$  are Bernoulli variables, K is geometrically distributed and we have

(2.7) 
$$E[K] = \frac{1}{P(I_1 = 1)}.$$

Let  $X_i(\varepsilon_i)$  denote the result on the jth test of the ith element:

(2.8)  $X_j(\varepsilon_i) = 1$  if the result on test j with element i is positive i if the result on test j with element i is negative.

Let  $S_r(\varepsilon_i) = \sum_{i=1}^r X_j(\varepsilon_i)$ . Suppose  $T_i = n$ , and  $S_{T_i} = m$ . Then (2.5) is equivalent to

(2.9) 
$$\frac{\alpha^{m}(1-\alpha)^{n-m}.\Pi}{\alpha^{m}(1-\alpha)^{n-m}.\Pi+\beta^{m}(1-\beta)^{n-m}.(1-\Pi)} \geqslant \Pi^{*}$$

or

(2.10) 
$$LR_n(\varepsilon_i) = \left(\frac{\alpha}{\beta}\right)^m \left(\frac{1-\alpha}{1-\beta}\right)^{n-m} \geqslant \frac{\Pi^*(1-\Pi)}{\Pi(1-\Pi^*)}.$$

 $LR_n(\varepsilon_i)$  is the likelihood ratio for  $\epsilon_i$  after n tests.

We can conclude now that whenever we have n trials on element  $\varepsilon_i$  with m positive results, so that (2.10) is satisfied, then there is no need for additional testing and i = K.

We can therefore identify the stopping and selection region as follows: stop and select  $\varepsilon_i$  whenever  $LR_n(\varepsilon_i) \ge \Pi^*(1-\Pi)/\Pi(1-\Pi^*)$ . That is,

$$\left\{T_i=n,\,I_i=1\right\}=\left\{LR_n(\varepsilon_i)\geqslant\frac{\Pi^*(1-\Pi)}{\Pi(1-\Pi^*)}\right\}.$$

It still remains to identify the stopping and rejection region. Consider the problem of testing the hypothesis  $H_0$ : " $\varepsilon_1$  is an A-element" versus the alternative  $H_1$ : " $\varepsilon_1$  is an  $\overline{A}$ -element."

Lemma 2.1. Using a Bayesian framework with  $P(H_0) = \Pi$ , we have that  $P_{H_1}(accepting \ H_0) = (\gamma/(1-\Pi))(1-\Pi^*)$  and  $P_{H_0}(rejecting \ H_0) = 1-(\Pi^*/\Pi)\gamma$  are equivalent to  $P(H_0|accepting \ H_0) = \Pi^*$  and  $P(accepting \ H_0) = \gamma$ .

PROOF.  $P(\text{accepting } H_0) = \prod P_{H_0}(\text{accepting } H_0) + (1 - \prod)P_{H_1}(\text{accepting } H_0)$  and

$$P(H_0|\text{accepting }H_0) = \frac{P(H_0) \cdot P_{H_0}(\text{accepting }H_0)}{P(\text{accepting }H_0)}.$$

Let  $(T_1, I_1)$  denote the variables for any stationary procedure in our original problems and let  $\gamma = P(I_1 = 1)$ . Consider a W.S.P.R.T. with  $P(\text{type } I \text{ error}) = 1 - (\Pi^*/\Pi)\gamma$  and  $P(\text{type } II \text{ error}) = \gamma(1 - \Pi^*/1 - \Pi)$ . Applying Lemma 2.1 to this test, we have  $P(H_0/\text{accepting } H_0) \ge \Pi^*$  and  $P(\text{rejecting } H_0) = \gamma$ . Let  $T^*$  be the W.S.P.R.T. stopping time. Then we have  $E[T^*] \le E[T]$  and so  $E[T^*]/\gamma \le E[T]/\gamma$ . We can conclude that for any stationary procedure that satisfies (2.9) there is a W.S.P.R.T. which satisfies (2.9) and which does at least as well in terms of expected cost.

The W.S.P.R.T. is defined by two constants, say  $B_0$ ,  $B_1$ ,  $B_0 \le 1 < B_1$ ; sampling is continued so long as

$$(2.11) B_0 \le LR_n < B_1.$$

We have already identified the upper bound as

(2.12) 
$$B_1 = \frac{\Pi^*(1-\Pi)}{\Pi(1-\Pi^*)}.$$

To identify  $B_0$  we need two lemmas.

LEMMA 2.2. For the W.S.P.R.T.,  $E_{H_0}[T]$ ,  $E_{H_1}[T]$ , and E[T] are nonincreasing functions of  $B_0$ .

PROOF. Immediate.

LEMMA 2.3. For the W.S.P.R.T.,  $P(accepting H_0|LR_n)$  is a nondecreasing function of  $LR_n$ .

PROOF. Note first that we have defined  $LR_n$  as the likelihood under  $H_0$  divided by the likelihood under  $H_1$ , and that the acceptance region of  $H_0$  was  $\{LR_n \ge B_1\}$ . Given  $LR_n$ , the continuation region can be viewed as a new W.S.R.R.T. with boundaries  $B_0/LR_n$ ,  $B_1/LR_n$  for observations n+1, n+2,  $\cdots$ . Since both new boundaries are decreasing functions of  $LR_n$ , the probability of exiting through the upper boundary is an increasing function of  $LR_n$ . For  $LR_n < B_0$  or  $LR_n \ge B_1$  the result is immediate.

THEOREM 2.1. Among all procedures satisfying (2.5) the W.S.P.R.T. with  $B_0 = 1$  and  $B_1 = (\Pi^*(1 - \Pi)/\Pi(1 - \Pi^*))$  minimizes  $E[T]/P(\text{selecting } \epsilon_1)$ .

PROOF. Follows from Lemmas 2.1, 2.2 and 2.3.

**3.** N > 2. For N > 2 we have  $(T_{11}, I_{11}), (T_{12}, I_{12}), \cdots, (T_{1K_1}, I_{1K_1}); (T_{21}, I_{21}), (T_{22}, I_{22}), \cdots, (T_{2K_2}, I_{2K_2}); (T_{N_1}, I_{N_1}), (T_{N_2}, I_{N_2}), \cdots (T_{NK_N}, I_{NK_N}).$  Our expected cost is  $E[C \cdot \sum_{m=1}^{N} \sum_{j=1}^{K} T_{mj}].$ 

Our problem is to minimize the expected cost subject to (2.5) for each of the N elements. Since N is a constant, we have an N-times repeated problem of choosing one element at a time, minimizing  $E[\sum_{j=1}^{K} T_{mj}]$  subject to (2.5). If we use a weaker criterion, namely that (2.5) is to be satisfied on the average, then due to overshoot over the boundaries we can economize and adjust the upper boundary  $B_1$  according to the accumulated average odds.

4. Remarks. It is clear from the analysis that the results are general enough to include the case for which the test response is a random variable with a distribution which depends on  $\overline{A}$  and on  $\overline{A}$ . If tests are not conditionally independent the theory may fail.

## REFERENCES

[1] FISHER, R. A. (1952). Sequential experimentation. Biometrics 8 183-187.

DEPARTMENT OF STATISTICS THE HEBREW UNIVERSITY JERUSALEM, ISRAEL