Introductory Sequential Methodologies with Applications: An Appreciation

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**!!!** Greetings from Connecticut **!!!** 

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1. Introduction and Key Ingredients:

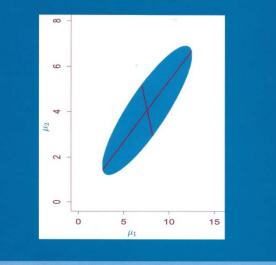
- Statistical tests or confidence interval estimation with a predetermined sense of accuracy
- Some sense of optimality in place
- Appropriate sample size determination
- Appropriate data collection strategies
- Gather data sequentially, step-by-step, and "learn" as we proceed

Some Key References:

- Mahalanobis (1940)
- Stein (1945,1949)
- Wald (Wiley: 1947)
- Mukhopadhyay and Solanky (Dekker: 1994, Chapter 2)

- Ghosh, Mukhopadhyay, and Sen (Wiley: 1997, Chapter 6)
- Mukhopadhyay, Datta, and Chattopadhyay, eds. (Dekker: 2004)
- Mukhopadhyay and de Silva (CRC: 2009)
- Note 1: The last-mentioned book includes a large number of standard methodologies with accompanying computer programs in executable forms
- Note 2: We will implement some of them in a number illustrations to follow

Sequential Methods and Their Applications



#### Nitis Mukhopadhyay Basil M. de Silva

CRC Press Taylor & Francis Group

## 2. Sequential Probability Ratio Test (SPRT)

X: A response variable with its probability distribution  $f(x;\theta)$ X may be discrete or continuous,  $\theta$  in  $\Theta$  unknown We may want to test

$$\begin{split} H_0: \theta &= \theta_0 \text{ versus } H_a: \theta = \theta_1, \, \theta_1 > \theta_0 \text{ and } \theta_0, \theta_1 \text{ from } \Theta. \\ \text{With } n \text{ independent responses } x_1, x_2, ..., x_n, \text{ Neyman-Pearson} \\ \text{theory will provide the most powerful (MP) test: Reject } H_0 \text{ iff} \\ \frac{\prod_{i=1}^n f(x_i; \theta_1)}{\prod_{i=1}^n f(x_i; \theta_0)} \geq k, \text{ that is the ratio is "large".} \\ \text{The cut-off } k(>0) \text{ is chosen by the fixed size of the test, } \alpha. \\ \text{We may also require power } 1 - \beta \text{ with preassigned } \alpha, \beta. \text{ Then?} \end{split}$$

#### Let us look at a "Normal Mean" example.

#### 2.1. Normal Mean Example

 $X_1, ..., X_n$  are random samples from  $N(\theta, \sigma^2)$  with  $\theta$  unknown, but  $\sigma^2$  known. We want to test

 $H_0: \theta = \theta_0$  versus  $H_a: \theta = \theta_1, \ \theta_1 > \theta_0.$ 

Detect effect size " $\theta_1 - \theta_0$ " efficiently!

The N-P MP level  $\alpha$  test would reject  $H_0$  iff  $\overline{X}$  is "large",

that is iff  $\overline{X} > \theta_0 + z_{\alpha} \sigma / \sqrt{n}$ 

where  $z_{\alpha}$  is the upper 100 $\alpha$ % point of N(0, 1) distribution. We further demand that type-II error must be  $\leq \beta$ , then the optimal fixed-sample-size n must be determined as:

$$n = \left(rac{\left(z_{lpha} + z_{eta}
ight)\sigma}{ heta_1 - heta_0}
ight)^2 \equiv n^*, ext{ say.}$$

SPRT will provide much more economical *n*.

#### 2.2. Wald's SPRT for General Distribution f

Having fixed  $\alpha, \beta$  we determine  $A = \frac{1-\beta}{\alpha}, B = \frac{\beta}{1-\alpha}$ . Then, keep observing  $X_1, X_2, \ldots$  one by one in a sequential manner until the likelihood ratio

$$\Lambda_{j} = \frac{\prod_{i=1}^{j} f\left(x_{i}; \theta_{1}\right)}{\prod_{i=1}^{j} f\left(x_{i}; \theta_{0}\right)}$$

goes outside the interval (B, A) first time.

Here, "A" indicates "large" and "B" indicates "small".

Then, we determine required number of observations N as:

 $N \equiv \text{first integer } j(\geq 1) \text{ such that } \Lambda_j \geq A \text{ or } \Lambda_j \leq B;$ 

We will accept  $H_a$  if  $\Lambda_N \ge A$ ; accept  $H_0$  if  $\Lambda_N \le B$ .

Although  $P(N < \infty) = 1$  w.p.1, N ought to be truncated if it asks to go beyond  $n^*$  observations in the normal example.

### 2.3. Optimality

Wald and Wolfowitz (1948) proved the following remarkable optimality property of SPRT among all tests (including the **N-P MP** test) with comparable  $\alpha, \beta$  (included in Mukhopadhyay and de Silva 2009, p. 43).

Optimal property of the SPRT: Consider testing  $H_0: \theta = \theta_0$ versus  $H_a: \theta = \theta_1$ . Among all tests with type-I error probability  $\leq \alpha$ , type-II error probability  $\leq \beta$ ,  $\alpha + \beta \leq 1$ , with  $E_{\theta}[N] < \infty$ when  $\theta = \theta_0, \theta_1$ , Wald's SPRT with error probabilities  $\alpha, \beta$  has the minimum  $E_{\theta}[N]$  when  $\theta = \theta_0, \theta_1$ .

#### 2.4. Truncation of the General SPRT

Suppose we can observe at most K observations only. Then we will implement the **truncated SPRT** as follows:

Keep observing  $X_1, X_2, ...$  one by one in a sequential manner checking whether the corresponding  $\Lambda_j$  goes outside (B, A)according to sampling stategy. If a decision is made with Nsmaller than K, then N is te sample size. if no decision is made even when the  $K^{\text{th}}$  response is observed, then we stop at N = K: Decide to accept  $H_0(H_a)$  if  $\Lambda_K \leq (>)\frac{1}{2}(A+B)$ .

In the normal problem, we should pick  $K = n^*$  given earlier!

#### 2.5. Back to Normal Distribution Illustration

Illustration Normal with  $\theta$  and  $\sigma^2$ 

$$\Lambda_{n} = \exp\left\{-(\theta_{0} - \theta_{1})\sigma^{-2}\sum_{i=1}^{n}X_{i} - n(\theta_{1}^{2} - \theta_{0}^{2})\sigma^{-2}/2\right\}$$

$$N \equiv \text{first } n(\geq 1) \text{ such that } \sum_{i=1}^{n}X_{i} \geq c_{1}\ln A + nc_{2},$$
or  $\sum_{i=1}^{n}X_{i} \leq c_{1}\ln B + nc_{2},$  with  $c_{1} = \sigma^{2}(\theta_{1} - \theta_{0})^{-1},$ 
 $c_{2} = (\theta_{0} + \theta_{1})/2.$ 
Accept  $H_{a}(H_{0})$  if  $\sum_{i=1}^{n}X_{i} \geq c_{1}\ln A + nc_{2}(\leq c_{1}\ln B + nc_{2}).$ 

Data Simulations with 10,000 Replication

$$\theta_0 = 1, \ \theta_1 = 2, \ \alpha = 0.05, \ \beta = 0.1$$
  
 $H_0: \theta = 1 \text{ versus } H_a: \theta = 2$   
 $\theta_0 = 1, \ \theta_1 = 2, \ \alpha = 0.05, \ \beta = 0.1$   
with  $\sigma^2 = 4, \ n^* = 34.27;$   
with  $\sigma^2 = 16, \ n^* = 137.08.$ 

Implement Seq03.exe from Mukhopadhyay and de Silva (2009)

In what follows, I take liberty in showing some screen shots in order to illustrate how a job runs.

Select one of the following programs:

1	SimAB:	Simulation of alpha + beta for a test
2	SPRT:	SPRT for given or simulated data
3	OC:	Computation of OC and ASN functions
0	Exit	

Enter your choice :2

SPRT for H0:theta - theta0 versus H1:theta - theta1

****	****	****	********	*****	***
*					*
*	ID#	DISTRIBUTION	Theta	Known	*
*			(Parameter1)	Parameter2	*
*					*
*	1	Normal	mean	variance	*
*	2	Gamnia	scale	shape	*
*	З	Weibull	scale	shape	*
*	4	Erlang	exp. mean	k(integer)	*
*	5	Exponential	mean		*
*	6	Poisson	mean		*
*	7	Bernoulli	P(Succ	ess)	*
*	8	Geometric	P(Succ	ess)	٠
*	9	Negative Binomia	al mean	k	*
*					*
****	*****	*************	**********	**********	***

Input distribution ID# : 1

```
Enter theta0 = 1
theta1 = 2
alpha = 0.05
beta = 0.1
```

Do you want to change the input parameters (Y or N)? n

```
Would you like to truncate the SPRT when sample size reaches K (Y or N)? n
```

```
Do you want to simulate data or input real data?
Type S (Simulation) or D (Real Data) : s
```

Do you want to store simulated sample sizes (Y or N)? n

Number of replications for a simulation? 10000

```
Enter a positive integer(<10) to initialize
the random number sequence : 8
```

```
For the data simulation, input
mean of the normal distribution: 1
variance: 4
```

SPRT: H0:theta = 1.00 versus H1:theta = 2.00 at alpha =0.05 and beta =0.10 where theta is the mean of normal distribution with variance = 4.0

> Results for simulated data from Normal distribution with mean = 1.00

Number of Simulations Number of times H0 is accepted	=	10000 9611
Average sample size (n bar) Std. dev. of n (s)	=	18.75 14.18
Minimum sample size (min n)	=	2
Maximum sample size (max n)	=	130
**********************************	***	******

Table 1. Simulated results with 10,000 replications for untruncated SPRT of  $H_0: \theta = 1$  vs.  $H_a: \theta = 2$  for  $N(\theta, \sigma^2)$  with  $\sigma^2 = (4, 16)$  known, preassigned  $\alpha = 0.05, \beta = 0.1$ , true  $\theta = (1, 1.5, 2)$ .

$\sigma^2$	true $\theta$	$n^*$	$\overline{n}$	$s_n$	$p_{H_0}\%$	M in	Max	$Q_1$	$Q_2$	$Q_3$
4	1	34.27	18.75	0.142	96.11	2	130	9	15	24
	1.5	34.27	32.91	0.267	55.89	2	297	14	25	44
	2	34.27	22.43	0.154	7.30	2	147	12	18	29
16	1	137.08	69.52	0.549	95.66	6	571	32	53	90
	1.5	137.08	116.25	0.957	55.75	7	825	49	87	153
	2	137.08	81.86	0.568	9.04	7	526	41	66	106

Table 2. Simulated results with 10,000 replications for truncated SPRT test of  $H_0: \theta = 1$  vs.  $H_a: \theta = 2$  for  $N(\theta, \sigma^2)$  with  $\sigma^2 = (4, 16)$  known, preassigned  $\alpha = 0.05, \beta = 0.1$ , true  $\theta = (1, 1.5, 2)$ . Truncation at K.

$\sigma^2$	true $\theta$	K	$\overline{n}$	$s_{\overline{n}}$	Min	Max	$Q_1$	$Q_2$	$Q_3$	рн. %	$n_T$	$p_{H_0}_T$
4	1	35	17.15	0.099	2	35	9	15	24	93.79	1146	72.69
	1.5	35	23.72	0.107	2	35	14	25	35	55.77	3368	49.73
	2	35	20.00	0.098	2	35	12	18	29	10.84	1522	25.95
16	1	138	63.78	0.393	6	138	31	53	91	93.51	1017	71.98
	1.5	138	88.05	0.427	6	138	49	87	138	56.17	2916	51.13
	2	138	74.25	0.385	8	138	42	66	105	11.71	1377	26.94

The **Operations Characteristic** (OC) function:

 $L(\theta) =$ Probability of accepting  $H_0$  when  $\theta$  is true,

$$L(\theta_0) = 1 - \alpha$$
, and  $L(\theta_1) = \beta$ .

The Average Sample Number (ASN) function:

 $E_{\theta}[N]$  when  $\theta$  is true

Next, we show some plots for the OC function  $[(\theta, L(\theta))]$ and ASN function  $[(\theta, E_{\theta}[N])]$ :

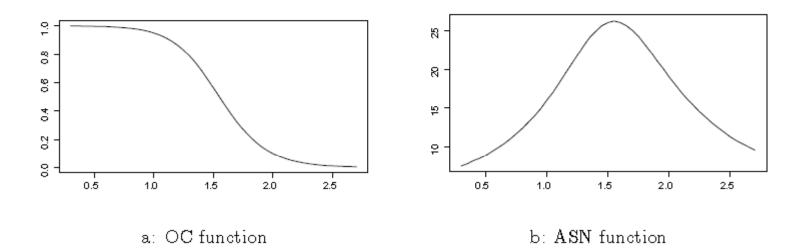


Fig. 1:  $H_0: \theta = 1$  vs.  $H_a: \theta = 2$  for  $N(\theta, 16), \alpha = 0.05, \beta = 0.1$ 

#### Real Data Illustration:

"Bodyfat" dataset from Statlib (lib.stat.cmu.edu/datasets/), submitted by Johnson (1996). Data include percentage of body fat determined by underwater weighing and various body circumference measurements for 252 men. We llooked at response variable (X): "Density determined from underwater weighing".

Some descriptive statistics of density\_underwater are shown:

We treated this dataset (X) as our population:  $N(\theta, 0.019^2)$ 

We set out to test:  $H_0: \theta = 1.055$  vs  $H_a: \theta = 1.060$  with  $\alpha = 0.05, \beta = 0.1$ , and  $K = 124(=n^*)$ . The SPRT terminated with N = 24 observations, and our decision was to accept  $H_0$  upon termination.

#### How about SPRT's in non-normal distributions?

In an extended version of the draft paper (with Yan Zhuang, expected Ph.D. August 2018), we have illustrations with simulations real datasets from

- Gamma (unknown scale, known shape): Body dimensions (Heinz et al., 2003),
- Lognormal (location unknown, scale known): Rosner's FEV (maximum forced expiratory volume in one second) data.

# 2.6. Remarks: Sequential Tests Under Unknown Nuisance Parameters and Other Situations:

Test on normal with unknown variance: Use program Seq04.exe (Mukhopadhyay and de Silva, 2009): Sequential *t*-test or Chi-square test or *F*-test ...

Discrete distributions (Binomial, Poisson, Negative binomial, Zero-inflated)

Unequal sampling at stages, perhaps according to some distribution (Mukhopadhyay and de Silva, 2005)

Two-sample or multi-sample tests

Scenarios where the response X may be multivariate ...

## 3. Fixed-Width Confidence Interval

### 3.1. Normal Mean Example

 $X_1, ..., X_n$  are independent, distributed as  $N(\theta, \sigma^2)$ , where  $\theta, \sigma^2$  are both unknown. Usual  $(1 - \alpha)$  confidence interval for  $\theta$  is:

$$I_n = \left[\overline{X}_n \pm t_{n-1,\alpha/2} \frac{S_n}{\sqrt{n}}\right],\,$$

where  $t_{n-1,\alpha/2}$  is the upper 50 $\alpha$ % point of the t-distribution with (n-1) degrees of freedom,  $\overline{X}_n$  is the sample mean, and  $S_n$  is the sample standard deviation. What should be n? Length of the confidence interval  $I_n$  is  $K_n = 2t_{n-1,\alpha/2} \frac{S_n}{\sqrt{n}}$ , which is random that might be (very) large for fixed n even if unknown  $\sigma$  is "small". That is,  $I_n$  may be too wide for practical use.

Therefore, it may be necessary to construct a confidence interval  $J_n$  for  $\theta$  which satisfies two requirements:

1)  $P\{\theta \in J_n\} \ge 1 - \alpha;$ 

2) Length of  $J_n \leq 2d, d(>0)$  is preassigned half-width. This  $J_n$  is called  $(1 - \alpha)$  fixed-width confidence interval for  $\theta$ . Such  $J_n$  can not be constructed by any fixed-sample-size procedure (Dantzig 1940) if n is predetermined and fixed. The fixed-width confidence interval  $J_n = [\overline{X}_n \pm d]$  for  $\theta$  also has preset  $(1 - \alpha)$  confidence if the required sample size nis determined as follows:

$$n\geq z_{lpha/2}^2\sigma^2/d^2\equiv n^*,$$
 say.

This  $n^*$  is called the **optimal fixed sample size** had  $\sigma^2$  been known. But, magnitude of  $n^*$  remains unknown! Some adaptive estimation methods for  $n^*$  are provided next.

## 3.1.1. Two-Stage Sampling

Mahalanobis (1940), Stein (1945,1949)

Begin with  $m(\geq 2)$  pilot observations  $X_1, ..., X_m$  and obtain the sample variance  $S_m^2$  from  $\{X_1, X_2, ..., X_m\}$  to estimate  $\sigma^2$ . Recall that  $n^*$  was  $z_{\alpha/2}^2 \sigma^2/d^2$ . Let the final sample size:  $N = \max\left\{m, \left\langle t_{m-1,\alpha/2}^2 S_m^2/d^2 \right\rangle + 1\right\},$ where  $\langle u \rangle$  = the largest integer  $\langle u$ . **Example 1:** Suppose m = 20, and  $t_{m-1,\alpha/2}^2 S_m^2/d^2 = 12.82$ . Then,  $N = \max\{20, < 12.82 > +1\} = 20 \Rightarrow N = 20$ . **Example 2:** Suppose m = 20, and  $t_{m-1,\alpha/2}^2 S_m^2/d^2 = 63.27$ . Then,  $N = \max\{20, < 63.27 > +1\} = 64 \Rightarrow N = 64$ .

Generally speaking, we proceed as follows:

- a) if N = m, then we already have enough data in pilot stage. In this case, final dataset is  $\{X_1, X_2, ..., X_m\}$ ;
- b) if N > m, then we have too few data in pilot stage. Then, we gather (N m) new observations in the second stage.
  In this case, final dataset is {X<sub>1</sub>,..., X<sub>m</sub>, X<sub>m+1</sub>,..., X<sub>N</sub>}.

Final fixed-width confidence interval for  $\theta$ :  $J_N = [\overline{X}_N \pm d]$ . where  $\overline{X}_N$  is the sample mean from final dataset  $\{X_1, ..., X_N\}$ .

Big Result (Consistency or Exact Consistency): For fixed  $\theta, \sigma, \alpha$  and d, one has:  $P_{\theta,\sigma} \{ \theta \in J_N = [\overline{X}_N \pm d] \} \ge 1 - \alpha$ .

This fundamental construction of adaptive sampling in two-steps changed everything after Wald's construction of the SPRT.

## Simulations were carried out using the program Seq06.exe from Mukhopadhyay and de Silva (2009)

	** 1	$\cdots$ $\cdots$ $\cdots$	epheadenb	with a	$\cdots$ , $m$	- 10	
$\alpha$	d	$n^*$	$\overline{n}$	$s_{\overline{n}}$	$\overline{n}/n^*$	$\overline{p}$	$s_{\overline{p}}$
0.1	1.500	19.239	24.314	0.110	1.264	0.903	0.003
	1.000	43.289	54.272	0.252	1.254	0.906	0.003
	0.800	67.639	83.978	0.392	1.242	0.902	0.003
	0.600	120.246	149.965	0.703	1.247	0.904	0.003
	0.400	270.554	336.704	1.580	1.244	0.894	0.003
	0.200	1082.217	1346.401	6.318	1.244	0.898	0.003

Table 3a. Simulations of two-stage estimation strategy with 10,000 replications with  $\alpha = 0.1, m = 10$ 

	with 10,000 replications with $\alpha = 0.05, m = 10$										
$\alpha$	d	$n^*$	$\overline{n}$	$s_{\overline{n}}$	$\overline{n}/n^*$	$\overline{p}$	$s_{\overline{p}}$				
0.05	1.500	27.317	37.001	0.171	1.354	0.952	0.002				
	1.000	61.463	81.718	0.382	1.330	0.949	0.002				
	0.800	96.036	128.128	0.592	1.334	0.950	0.002				
	0.600	170.732	229.777	1.093	1.346	0.950	0.002				
	0.400	384.146	517.077	2.466	1.346	0.950	0.002				
	0.200	1536.584	2051.753	9.630	1.335	0.952	0.002				

Table 3b. Simulations of two-stage estimation strategy with 10,000 replications with  $\alpha = 0.05, m = 10$ 

Table 3c. Simulations results of two-stage estimation strategy with 10,000 replications with  $\alpha = 0.01, m = 10$ 

$\alpha$	d	$n^*$	$\overline{n}$	$S_{\overline{n}}$	$\overline{n}/n^*$	$\overline{p}$	$s_{\overline{p}}$
0.01	1.500	47.181	75.382	0.357	1.598	0.991	0.001
	1.000	106.158	169.157	0.795	1.593	0.989	0.001
	0.800	165.872	264.773	1.230	1.596	0.992	0.001
	0.600	294.884	470.074	2.222	1.594	0.990	0.001
	0.400	663.490	1062.446	5.022	1.601	0.990	0.001
	0.200	2653.959	4258.497	20.121	1.605	0.991	0.001

Stein's two-stage solution, though path-breaking, also came with some baggage of its own:

- a) Oversampling:  $E[N/n^*] > t_{m-1,\alpha/2}^2/z_{\alpha/2}^2(>1)$ . Not too disturbing!
- b) Asymptotic Oversampling: E[N/n\*] → t<sup>2</sup><sub>m-1,α/2</sub>/z<sup>2</sup><sub>α/2</sub>
  (> 1) as d → 0. This can be disturbing!
  Note that σ<sup>2</sup> was estimated only once through pilot data.
  Appropriate modifications were proposed and implemented
  by Mukhopadhyay (1980) and Mukhopadhyay and Duggan
  (1997) in order to gain substantial efficiency by drastically
  reducing the extent of oversampling via right choices of
  m under two-stage adaptive sampling methods.

Why not estimate  $\sigma^2$  sequentially, multiple times?

Table	e 4. Asymptotic Ov	ersampling Percentage								
$t_{m-1,\alpha/2}^2/z_{\alpha/2}^2$ values for $\alpha = 0.1, 0.05, 0.01,$										
m = (10, 20) in the Two-Stage Procedure										
	m = 10	m = 20								
$\alpha$	oversampling%	oversampling%								
0.10	1.2420 (24.20%)	1.1051(10.51%)								
0.05	1.3321 (33.21%)	1.1404(14.04%)								
0.01	1.5918 (59.18%)	1.2336(23.36%)								

#### Real Data Illustration:

We used the infection risk of hospital infection data from Kutner et al. (2005) on 113 hospitals in the US, 1975-76. We treated the 113 observed values of infection risk as population data. Normality checks went smoothly.

Table 5. Single run to estimate the mean of infection risk with  $\alpha = 0.05$ ,

d	$n^*$	n	$\overline{X}_n$	$S_n^2$	95% CI
0.6	19.02	25	4.26	1.57	(3.66, 4.86)
0.5	27.38	35	4.31	1.97	(3.81, 4.81)
0.4	42.79	55	4.40	1.97	(4.00, 4.80)

True mean infection risk 4.355 lies within each conf interval.

#### 3.1.2. Purely Sequential Sampling

# Anscombe (1952), Ray (1957), and Chow and Robbins (1965)

Begin with  $m(\geq 2)$  pilot observations  $X_1, ..., X_m$  and obtain one additional observation at-a-time giving rise to  $(n, \overline{X}_n, S_n^2)$ , successively for n = m, m + 1, ... as needed. Recall that  $n^*$  was  $z_{\alpha/2}^2 \sigma^2/d^2$ . Let the **final sample size**:  $N = \min \left\{ n \geq m : n \geq z_{\alpha/2}^2 S_n^2/d^2 \right\}$ . In this case, final dataset is  $\{X_1, ..., X_m, ..., X_N\}$ .

Final fixed-width confidence interval for  $\theta$ :  $J_N = [\overline{X}_N \pm d]$ . with  $\overline{X}_N$ , the sample mean from final dataset  $\{X_1, ..., X_N\}$ . **Properties:** 

 $P_{\theta,\sigma}(N < \infty) = 1.$ Asymptotic Consistency:  $\lim_{d \to 0} P_{\theta,\sigma} \{\theta \in J_N\} = 1 - \alpha.$ Asymptotic Efficiency:  $\lim_{d \to 0} E_{\theta,\sigma}[N/n^*] = 1.$ This methodology is also asymptotically second-order efficient (Ghosh and Mukhopadhyay 1981).

Next, we show some simulated results.

	strategy with 10,000 replications, $\alpha = 0.05, m = 10$										
$\alpha$	d	$n^*$	$\overline{n}$	$s_{\overline{n}}$	$\overline{n}/n^*$	$\overline{p}$	$s_{\overline{p}}$				
0.05	1.500	27.317	25.423	0.082	0.931	0.923	0.003				
	1.000	61.463	59.531	0.124	0.969	0.941	0.002				
	0.800	96.036	94.469	0.147	0.984	0.943	0.002				
	0.600	170.732	169.660	0.190	0.994	0.948	0.002				
	0.400	384.146	383.407	0.276	0.998	0.948	0.002				
	0.200	1536.584	1536.069	0.558	1.000	0.952	0.002				

Table 6. Simulated results of purely sequential estimation strategy with 10,000 replications,  $\alpha = 0.05, m = 10$ 

_	strategy with 10,000 replications, $\alpha = 0.05, m = 20$								
$\alpha$	d	$n^*$	$\overline{n}$	$s_{\overline{n}}$	$\overline{n}/n^*$	$\overline{p}$	$S\overline{p}$		
0.05	1.500	27.317	27.104	0.062	0.992	0.942	0.002		
	1.000	61.463	59.686	0.120	0.971	0.941	0.002		
	0.800	96.036	94.553	0.145	0.985	0.943	0.002		
	0.600	170.732	169.677	0.189	0.994	0.948	0.002		
	0.400	384.146	383.364	0.276	0.998	0.948	0.002		
	0.200	1536.584	1536.060	0.557	1.000	0.949	0.002		

Table 7. Simulated results of purely sequential estimation strategy with 10,000 replications,  $\alpha = 0.05, m = 20$ 

#### **Real Data Illustration:**

Table 8. Single run to estimate the mean infection risk with  $\alpha = 0.05, m = 10$ for purely sequential procedure

	-	~	-	-	
d	$n^*$	N	$\overline{X}_N$	$S_n^2$	95% CI
0.6	19.02	20	4.20	1.87	(3.60, 4.80)
0.5	27.38	24	4.20	1.55	(3.70, 4.70)
0.4	42.79	48	4.46	1.94	(4.06, 4.86)

True mean infection risk 4.355 lies within each confidence interval.

3.2. Remarks: Sequential Fixed-Size Confidence Regions and Other Situations:

Discrete distributions (Binomial, Poisson, Negative binomial, Zero-inflated): Based on the MLE

Many other continuous distributions (Negative exponential, Exponential, Gamma, Lognormal ...): Based on the MLE

Multivariate and regression simultaneous confidence and point estimation problems (under other loss functions)

Unequal variances (treatment allocations)

Multiple comparisons ...

## Thank You

Any Question or Comment?