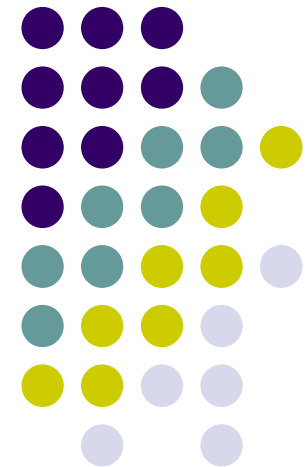


A comparison of interval estimates of the reliability coefficient obtained from Bayesian rejection sampling and the adjusted Searle method, in small studies

Ruth Pickering
Public Health Sciences and Medical Statistics

UNIVERSITY OF
Southampton
School of Medicine



The reliability coefficient (R)

in simple replication studies



Repeated readings of variable X taken on N subjects

$$X_{ik} = \mu + \alpha_i + \varepsilon_{ik}$$

subject $i = 1 \dots N$

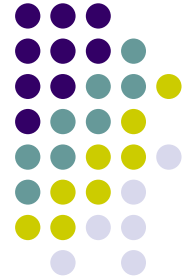
repeat $k = 1 \dots n_i$ - number of repeats can vary with subject

the α_i follow a Normal $(0, \sigma_{\text{subject}}^2)$ } independent
the ε_{ik} a Normal $(0, \sigma_{\text{repeat}}^2)$ }

$$R = \frac{\sigma_{\text{subject}}^2}{\sigma_{\text{subject}}^2 + \sigma_{\text{repeat}}^2} - \text{expressed as a \%}$$

R and the ICC are alternative names for the same thing

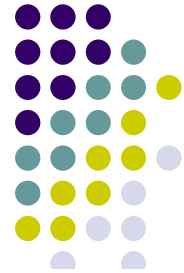
Rationale



- coverage of interval estimates for R depends on:
 - true value of R
 - N (subjects) & n (repeats)
 - distribution of α_i and ε_{ik}
- In small reliability studies
 - typically R = 60-100%, N=10-20, n=2-5
- SAS PROC MIXED
 - fits variance component models
 - doesn't directly produce estimates of R
 - includes an empirical Bayes method of rejection sampling giving credibility intervals for $\sigma_{\text{subject}}^2$, σ_{repeat}^2 and thus R
- Bayesian approach extends to more complex designs
 - eg inter-observer studies
- how does the Bayesian interval perform?

Confidence intervals (CI) for R

Searle's method for balanced data: $n_i=n$



One-way ANOVA for variable X including subject

The ratio of MS_{subject} and MS_{error} gives the F test

$$F = \frac{MS_{\text{subject}}}{MS_{\text{error}}}$$

F_L and F_U are the 2.5th and 97.5th percentiles of an

$F_{N(n-1)}^{N-1}$ Distribution

Searle's CI $\left\{ \frac{F/F_U - 1}{n + F/F_U - 1}, \frac{F/F_L - 1}{n + F/F_L - 1} \right\}$ is **EXACT**



Adjusted Searle's CI for unbalanced data

The fixed number of repeats, n , is replaced by:

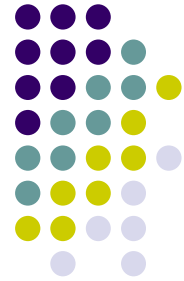
$$n_0 = \bar{n}_i - \frac{\text{Variance}(n_i)}{k \times \bar{n}_i},$$

when n_i varies over subjects to give the CI:

$$\left\{ \frac{F/F_U - 1}{n_0 + F/F_U - 1}, \frac{F/F_L - 1}{n_0 + F/F_L - 1} \right\}$$

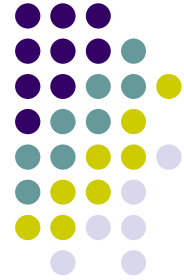
Note not exact if n_i varies

Bayesian intervals from SAS PROC MIXED



```
proc mixed data=datafile method=REML;  
class subject;  
model varX= ;  
random subject ;  
prior / nsample=10000 out=rejsamples;  
run;
```

- REML estimates of $\sigma^2_{\text{subject}}$ and σ^2_{repeat} requested
- subject included as a random effect
- doesn't directly produce an interval estimate for R
- `prior / nsample=10000 out=rejsamples;` requests 10,000 rejection samples be drawn and output to the file 'rejsamples'
- subsequent calculations on 'rejsamples' need to be requested by the user



Rejection sampling

- Bayesian monte-carlo technique
- samples from the joint posterior of $\sigma^2_{\text{subject}}$ and σ^2_{repeat}
- specifically developed for estimating variance components

Wolfinger RD & Kass RE. Nonconjugate Bayesian analysis of variance component models. *Biometrics* 2000; 56: 768-774.

- suggest 10,000 samples be drawn
- by default PROC MIXED uses Jeffrey's priors
 - non-informative
 - hyperparameters estimated from data



Motivating example

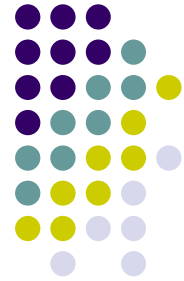
- **test-retest reliability of walking parameters in people with Parkinson's disease ≥ 6 months**
- **participants assessed twice**
 - a week apart (mean 6: min-max 3-14 days)
 - same time of day on each occasion,
 - no changes in medication
- **first invited 10 people, then doubled sample size to 20**
- **2 unwilling to travel to the laboratory (1 in each half)**
- **sporadic missing values**

EXAMPLE: Point (interval) estimate of R for walking parameters



parameter		# with 2 readings	Searle's	adjusted Searle's	Bayesian
Speed (meters/min)	first 9	8	74 (20, 94)	75 (21, 94)	75 (24, 94)
	all 18	17	79 (68, 95)	79 (52, 92)	79 (52, 92)
Stride Length (meters)	first 9	8	98 (90, 100)	98 (91, 100)	98 (91, 100)
	all 18	17	93 (83, 97)	93 (83, 97)	93 (83, 98)
Cadence (steps/min)	first 9	8	53 (-17, 88)	49 (-24, 86)	50 (6, 86)
	all 18	17	68 (32, 87)	67 (30, 86)	67 (31, 86)

SIMULATIONS TO COMPARE THE PROPERTIES OF THE ADJUSTED SEARLE AND BAYESIAN INTERVALS



- size of simulated datasets similar to example data
 - **N - number of subjects** - 10, 20, 30
 - **n - number of repeats** - 2, 3, 5
- balanced or unbalanced (50% with only 1-2 repeats)
- subject effects ~ normal(0,1)
or lognormal(0, 0.25) standardised to $\sigma_{\text{subject}}^2 = 1$
- error terms ~ Normal (0, σ_{repeat}^2)
- range of σ_{repeat}^2 giving R= 10%, 20%, ... , 80%, 90%
- 1000 datasets simulated for each combination
 - PROC MIXED simulated 10,000 rejection samples for each one

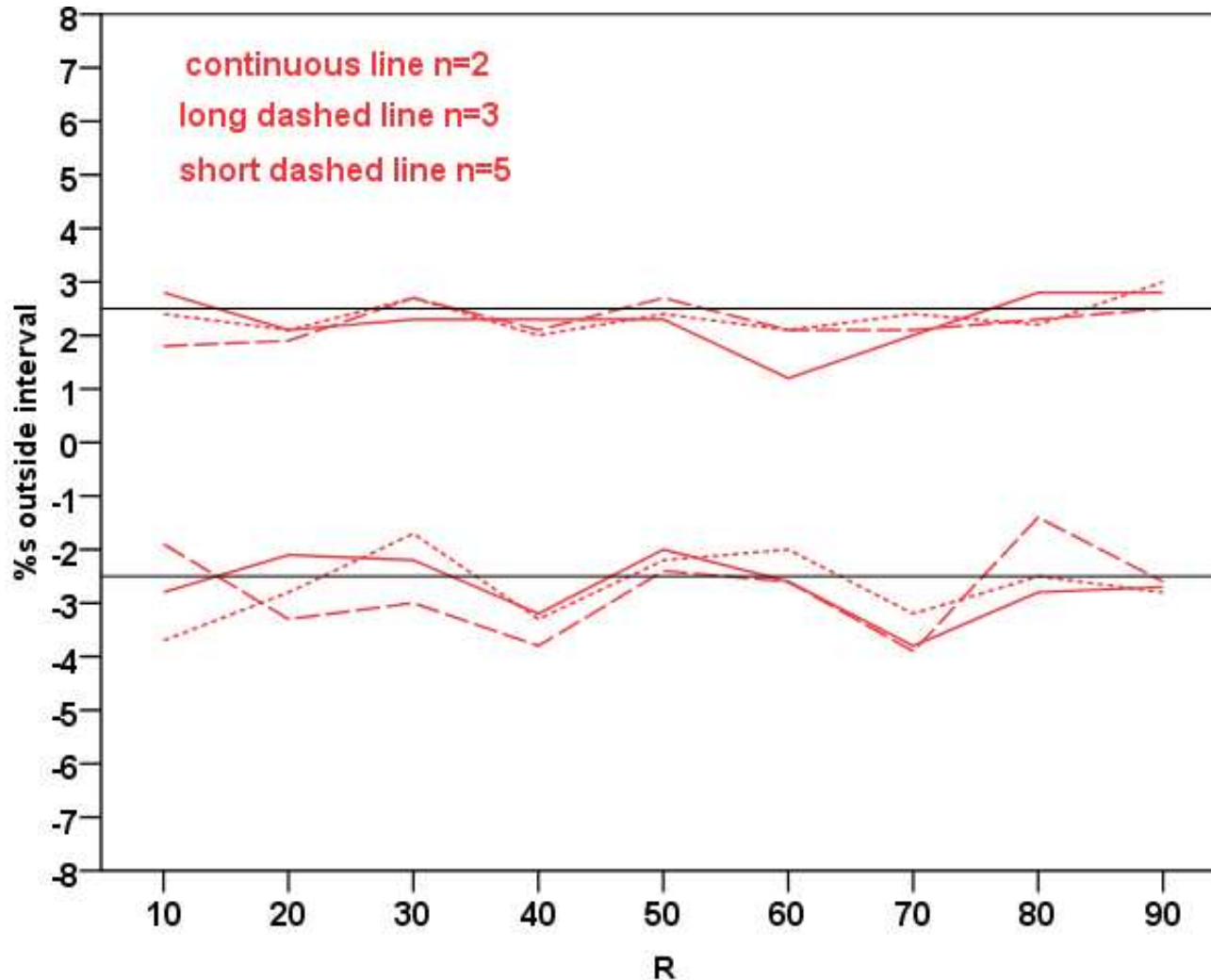
RESULTS - COVERAGE

upper lines - % true R > upper limit

lower lines - % true R < lower limit



Searle's CI with balanced data: number of subjects N=10



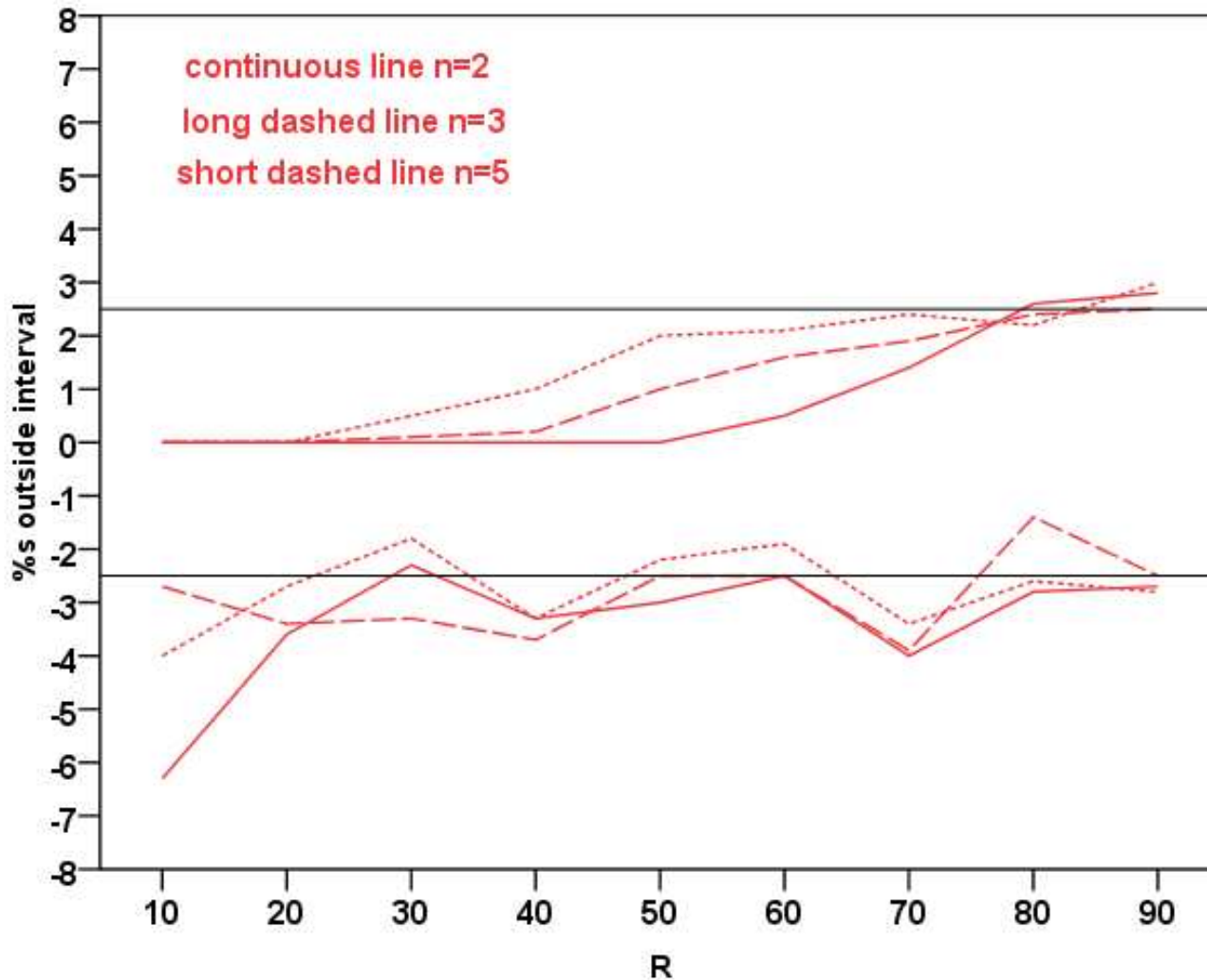
RESULTS - COVERAGE

upper lines - % true $R >$ upper limit

lower lines - % true $R <$ lower limit



Bayesian interval with balanced data: N=10

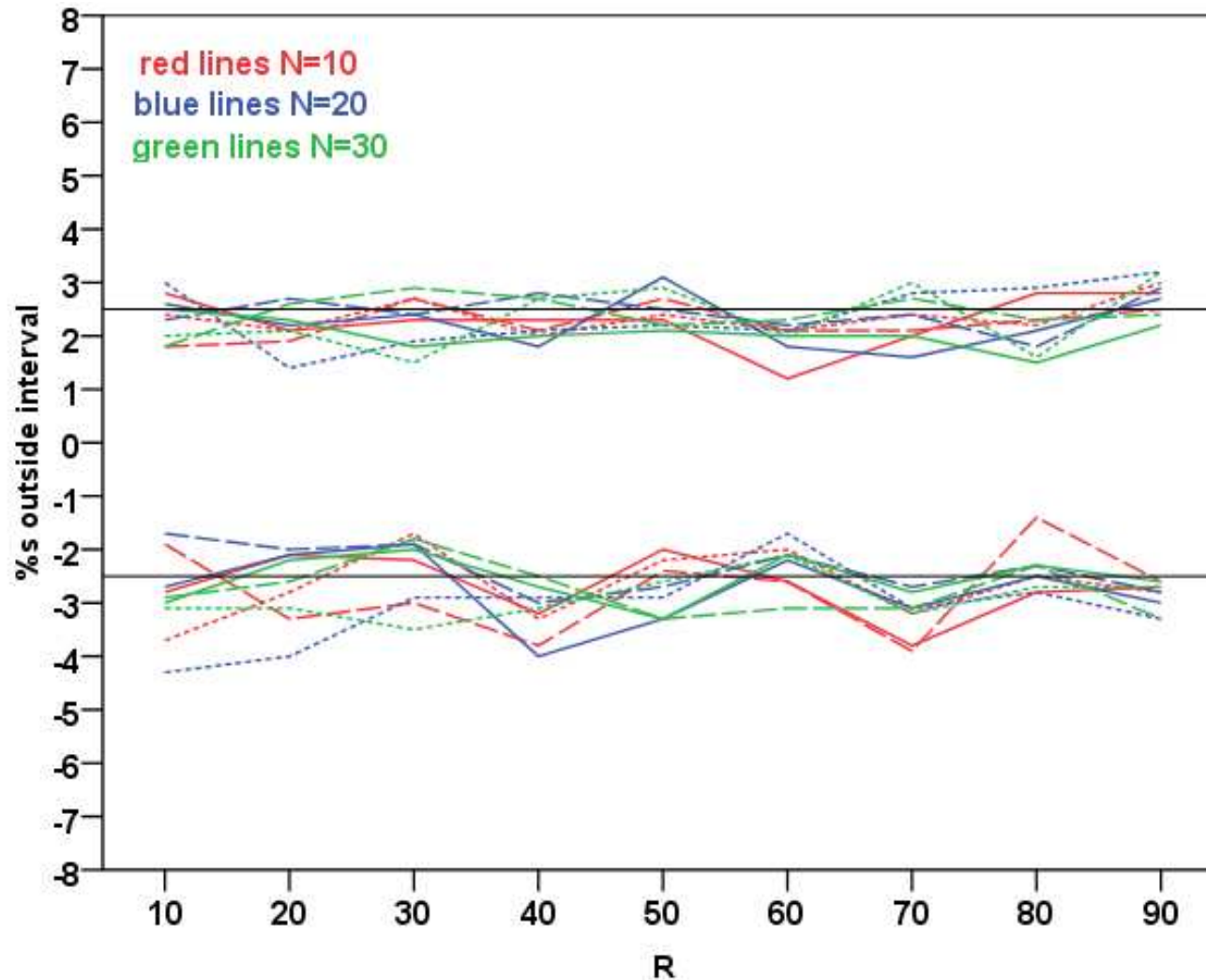


RESULTS - COVERAGE

upper lines - % true R > upper limit

lower lines - % true R < lower limit

Searle's CI with balanced normal data



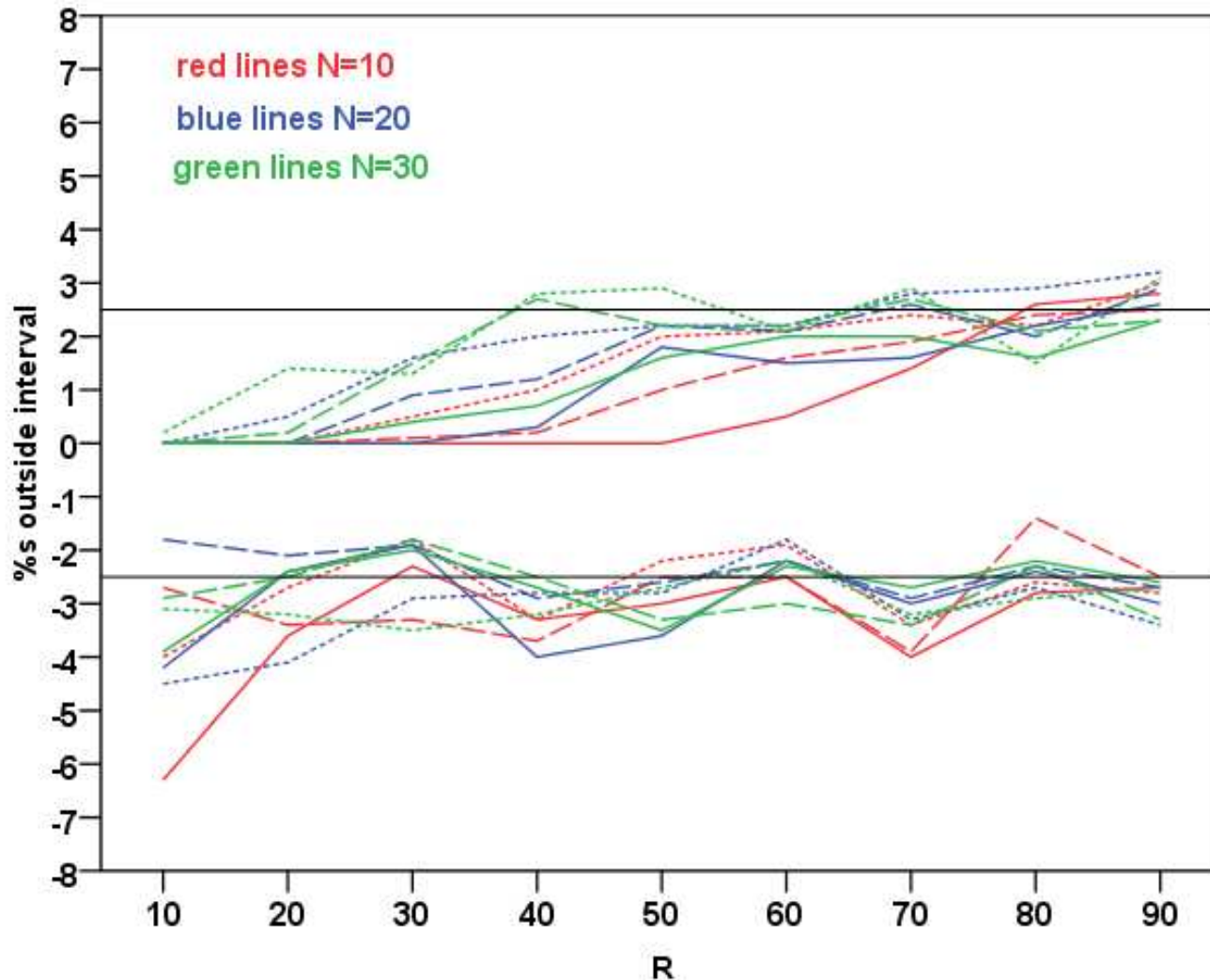
RESULTS - COVERAGE

upper lines - % true $R >$ upper limit

lower lines - % true $R <$ lower limit



Bayesian interval with balanced normal data

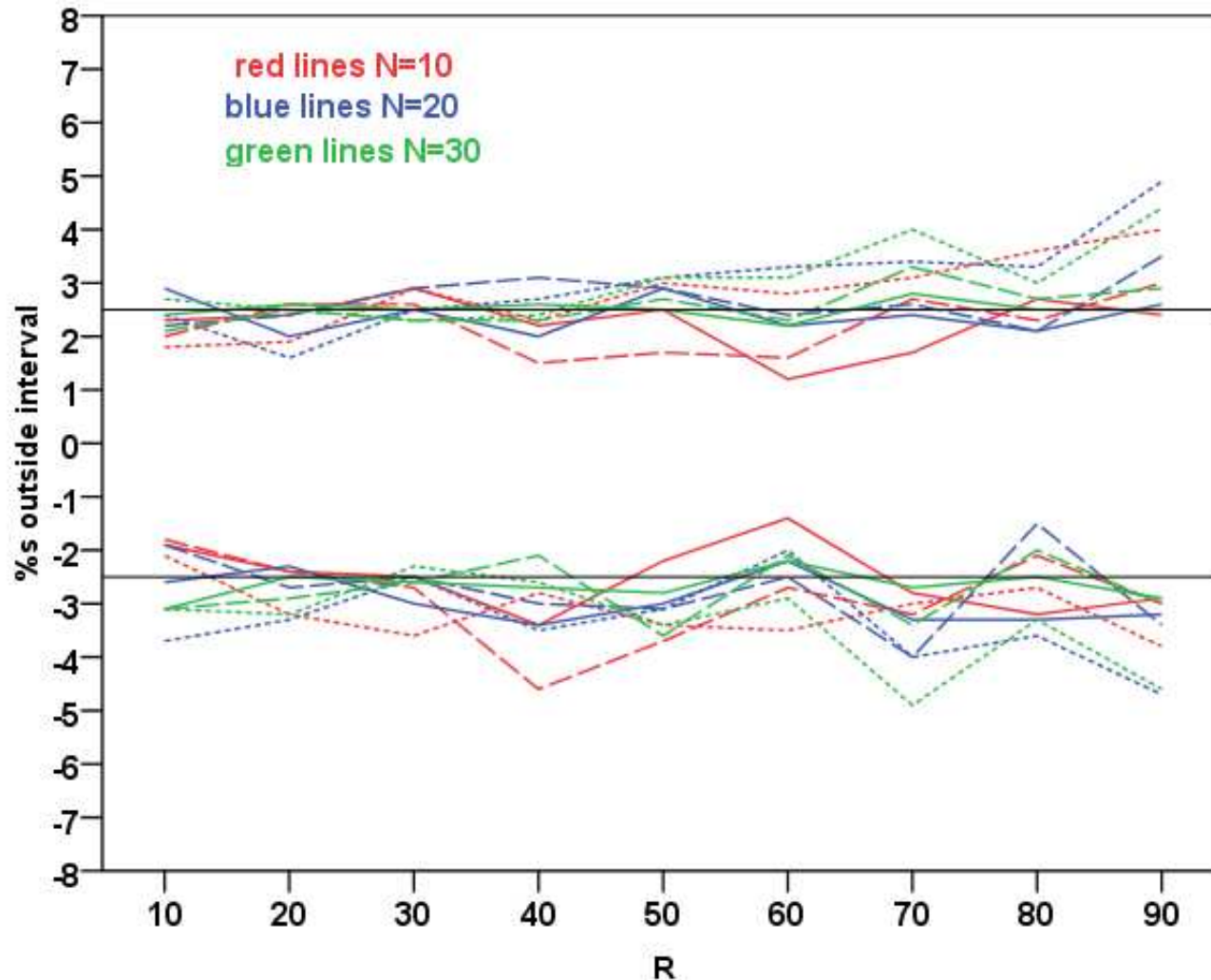


RESULTS - COVERAGE

upper lines - % true $R >$ upper limit

lower lines - % true $R <$ lower limit

Searle's CI with unbalanced normal data



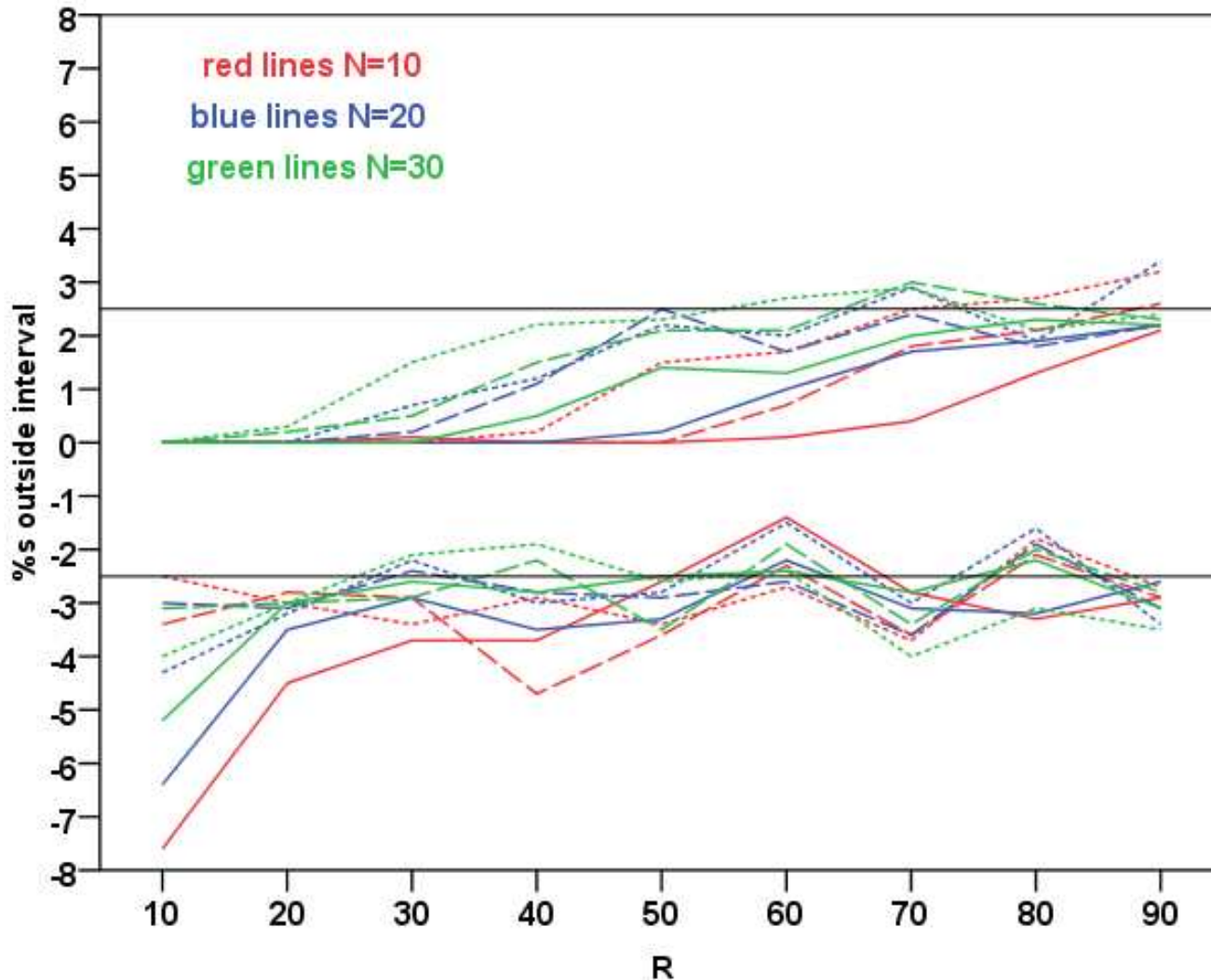
COVERAGE

lower lines - % true $R <$ lower limit

upper lines - % true $R >$ upper limit



Bayesian interval with unbalanced normal data



RESULTS - COVERAGE

upper lines - % true R > upper limit

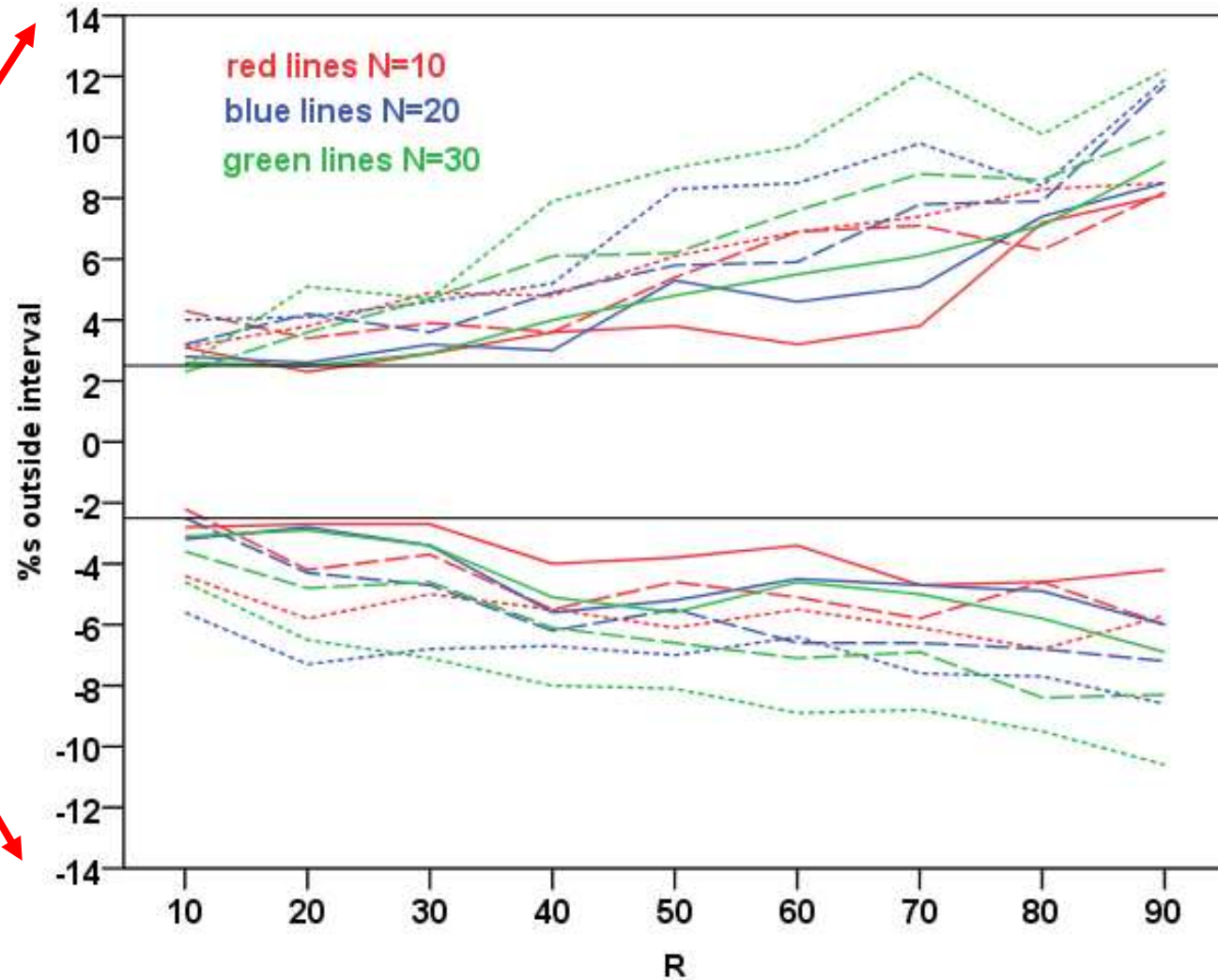
lower lines - % true R < lower limit



Searle's CI with balanced non-normal data

Note
change of
scale

14% vs 8%



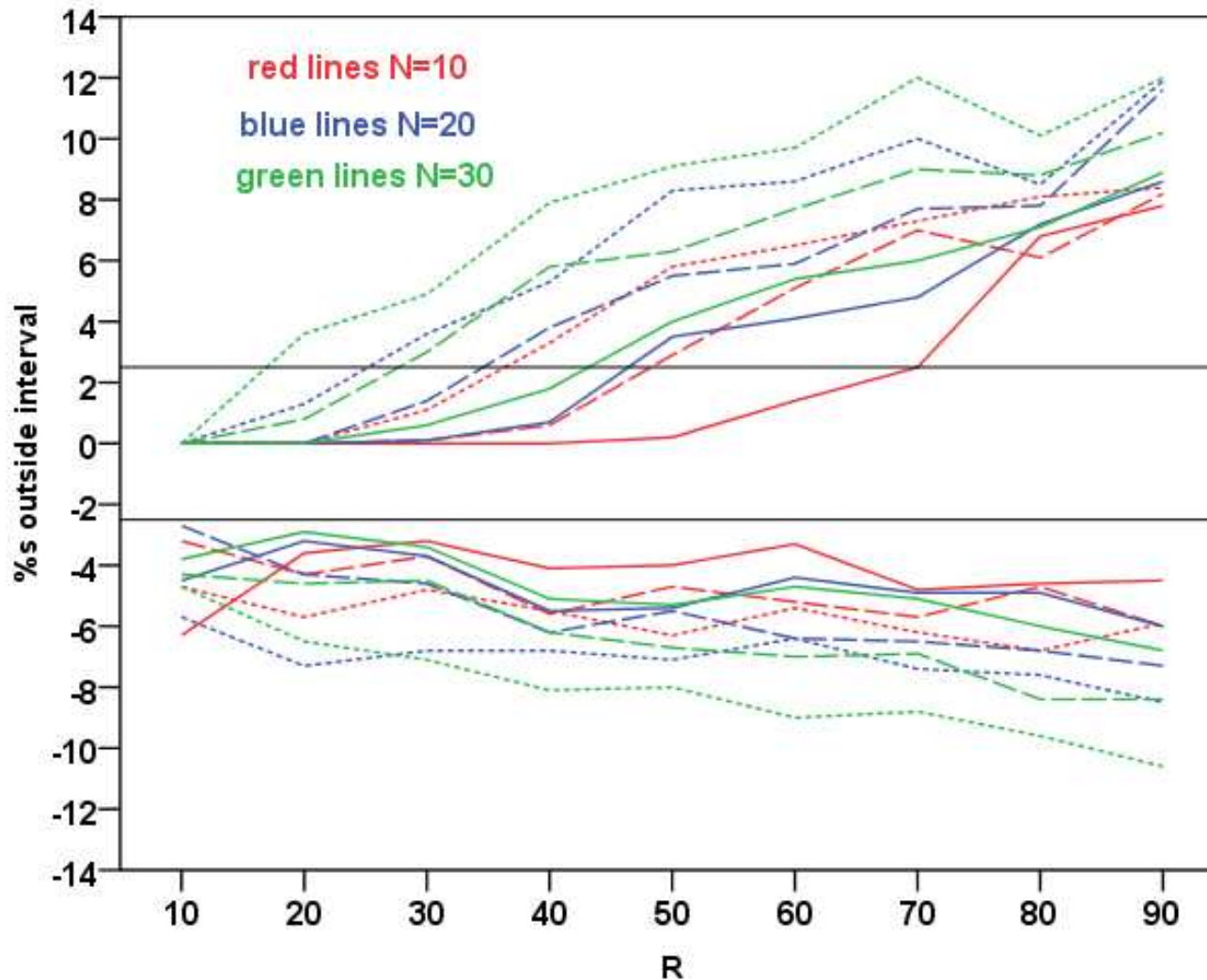
RESULTS - COVERAGE

upper lines - % true $R >$ upper limit

lower lines - % true $R <$ lower limit



Bayesian interval with balanced non-normal data





MEAN WIDTH OF INTERVALS

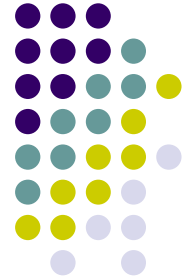
Normal balanced data

		Searle's		Bayesian
		mean width	mean +ve width	mean width
N=10 n=2	R=10	108.6	61.9	67.3
	R=90	34.4	34.2	33.8
N=10 n=5	R=10	52.3	44.3	47.0
	R=90	21.7	21.7	21.7
N=30 n=2	R=10	67.7	42.6	45.0
	R=90	15.4	15.4	15.4
N=30 n=5	R=10	29.0	25.7	25.9
	R=90	11.1	11.1	11.1

SUMMARY OF PROPERTIES OF INTERVALS



- **the Bayesian interval is too high for small R**
 - less of a problem for large N (subjects) & n (repeats)
 - less/no problem for R typical of reliability studies
 - evident in balanced/unbalanced, normal/non-normal data
- **Searle's and Bayesian intervals similar for large R in balanced normal datasets**
- **unbalanced data with large R: Searle's CI undercovers for large R, N & n: the Bayesian interval has close to nominal coverage for large R**
- **non-normal data: both intervals undercover for large R**
 - more of a problem for large N & n
- **Bayesian interval wider (up to 8%) when R small**
 - similar widths for large R typical of measurement studies



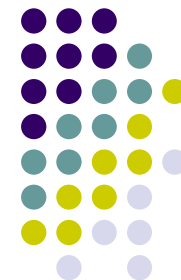
CONCLUSIONS

- **Bayesian interval easy to obtain from PROC MIXED with properties similar to Searle's CI for R typical of reliability studies**
 - probably not satisfactory for small R
- **balance no issue with the Bayesian interval**
- **serious under-coverage for both intervals at larger R when data is not normal**



REFERENCES

- Wolfinger RD & Kass RE. Nonconjugate Bayesian analysis of variance component models. *Biometrics* 2000; 56: 768-774.
- Donner A, Wells G. A comparison of confidence interval methods for the intraclass correlation coefficient. *Biometrics* 1985; 42: 401-412.
- Burdick RK, Graybill FA. Confidence intervals on variance components. Marcel Dekker: New York, 1992.
- Ukoumunne OC. A comparison of confidence interval methods for the intraclass correlation coefficient in cluster randomized trials. *Statistics in Medicine* 2002; 21: 3757-3774.
- Ukoumunne OC, Davison AC, Gulliford MC, Chinn S. Non-parametric bootstrap confidence intervals for the intraclass correlation coefficient. *Statistics in Medicine* 2003; 22; 3805-21.
- Littell RC, Milliken GA, Stroup WW, Wolfinger RD, Schabenberger O. *SAS[®] for Mixed Models*, second edition. SAS Publishing 2006.
- Searle SR. *Linear Models*. Wiley: New York, 1971.



Point estimates of R

- **Searle's method : based on estimates of $\sigma_{\text{subject}}^2$ and σ_{repeat}^2 incorporating n_0 , if n_i vary**
 - truncated at zero if $\sigma_{\text{subject}}^2$ -ve
- **R calculated from the REML estimates**
- **the median of the 10,000 R calculated from each rejection sample**

SUMMARY OF PROPERTIES OF POINT ESTIMATES



- **balanced normal datasets**
 - mean estimates ordered according to **true \approx Searle's $<$ truncated Searle's \approx REML $<$ median rejection**
 - for small N, n and R
 - eg N=10, n=2, R=10% \approx 10.3 $<$ 18.7 \approx 18.7 $<$ 30.5
 - all similar for larger R
 - eg N=10, n=2, R=90% \approx 87.7 \approx 87.7 \approx 87.7 \approx 87.8
- **unbalanced normal datasets**
 - similar pattern with greater over-estimation
 - eg N=10, n=2, R=10% \approx 10.2 $<$ 24.5 \approx 23.5 $<$ 39.6
 - eg N=10, n=2, R=90% \approx 87.7 \approx 87.7 \approx 87.5 \approx 87.4
- **non-normal datasets**
 - similar to balanced datasets
 - eg N=10, n=2, R=10% \approx 9.6 $<$ 18.5 \approx 18.5 $<$ 30.3
 - eg N=10, n=2, R=90% \approx 84.8 \approx 84.8 \approx 84.8 \approx 85.1



Use of point estimates

- **present R from REML estimates with the Bayesian interval?**

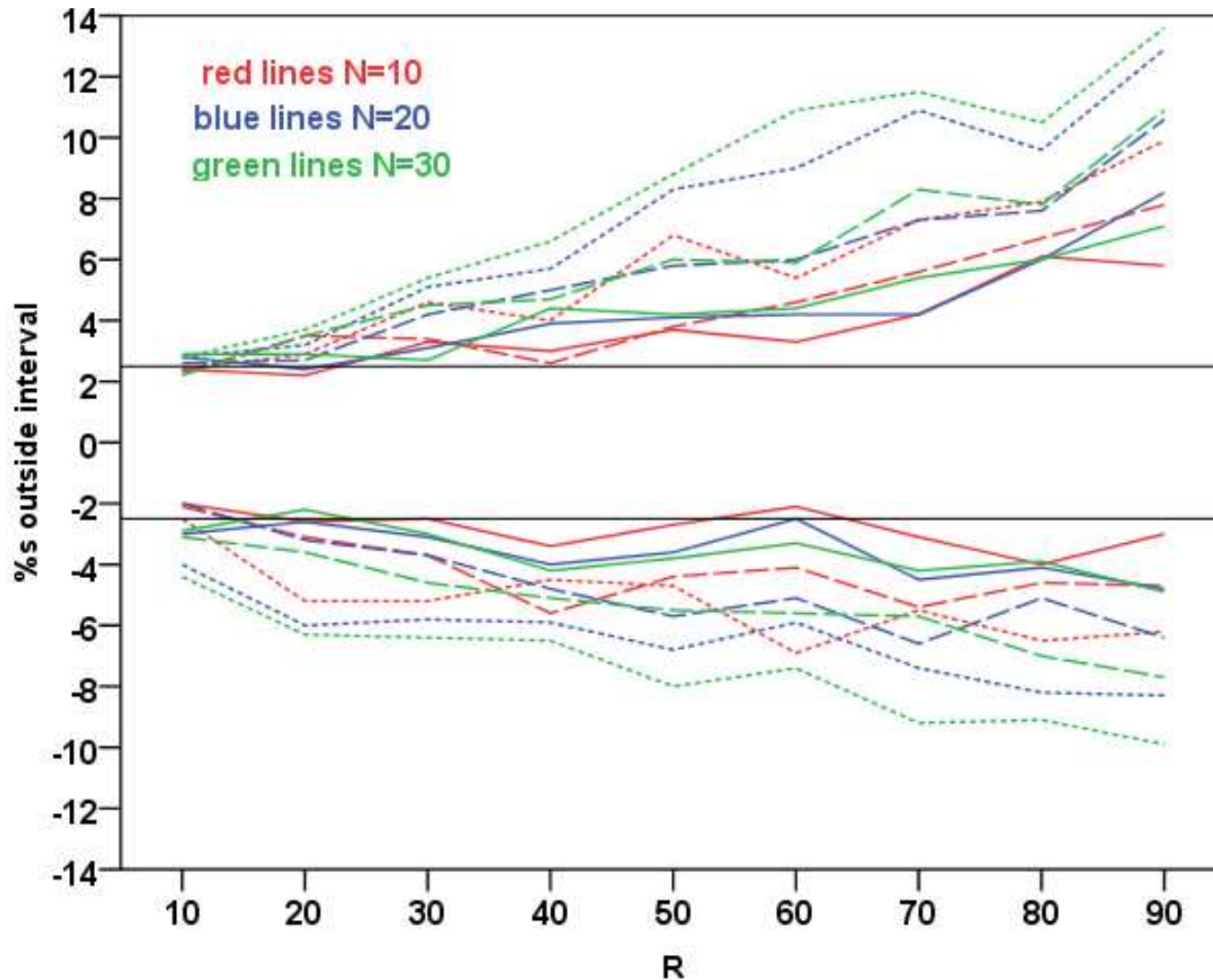
RESULTS - COVERAGE

upper lines - % true R > upper limit

lower lines - % true R < lower limit



Searle's CI with unbalanced non-normal data



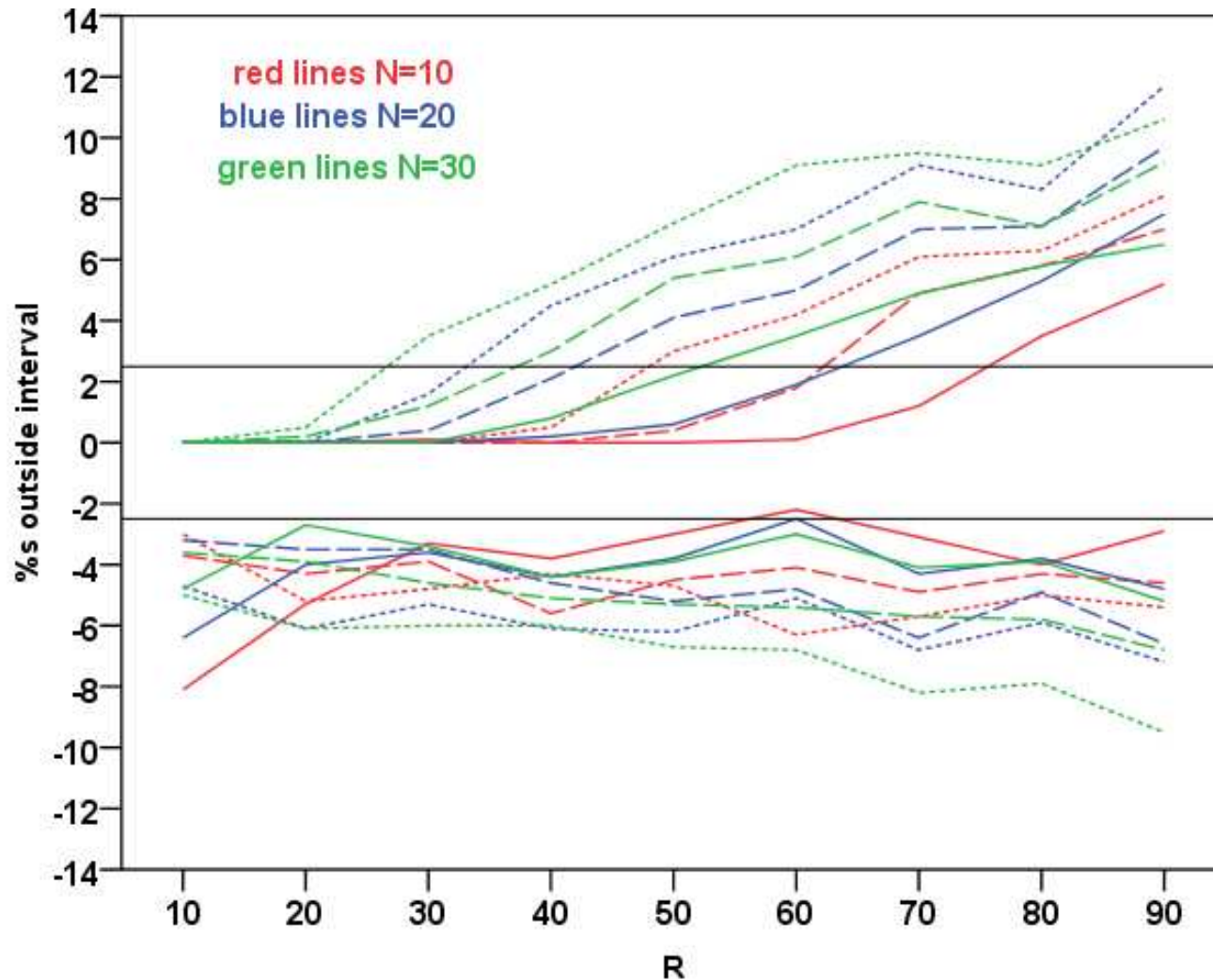
RESULTS - COVERAGE

upper lines - % true R > upper limit

lower lines - % true R < lower limit



Bayesian interval with unbalanced non-normal data



MEAN WIDTH OF INTERVALS

Normal unbalanced data



		Searle's		Bayesian
		mean width	mean +ve width	mean width
N=10 n=2	R=10	181.9	70.6	75.5
	R=90	91.6	53.7	48.7
N=10 n=5	R=10	69.5	50.7	57.1
	R=90	25.2	25.2	25.6
N=30 n=2	R=10	108.8	51.4	54.8
	R=90	21.7	21.7	21.9
N=30 n=5	R=10	40.9	32.2	33.6
	R=90	12.4	12.4	12.5



MEAN WIDTH OF INTERVALS

non-normal balanced data

		Searle's		Bayesian
		mean width	mean +ve width	mean width
N=10 n=2	R=10	108.5	61.4	67.0
	R=90	39.5	38.9	38.0
N=10 n=5	R=10	51.7	43.3	46.4
	R=90	24.9	24.9	25.0
N=30 n=2	R=10	67.7	42.4	44.8
	R=90	16.8	16.8	16.8
N=30 n=5	R=10	28.8	25.3	25.6
	R=90	12.0	12.0	12.0

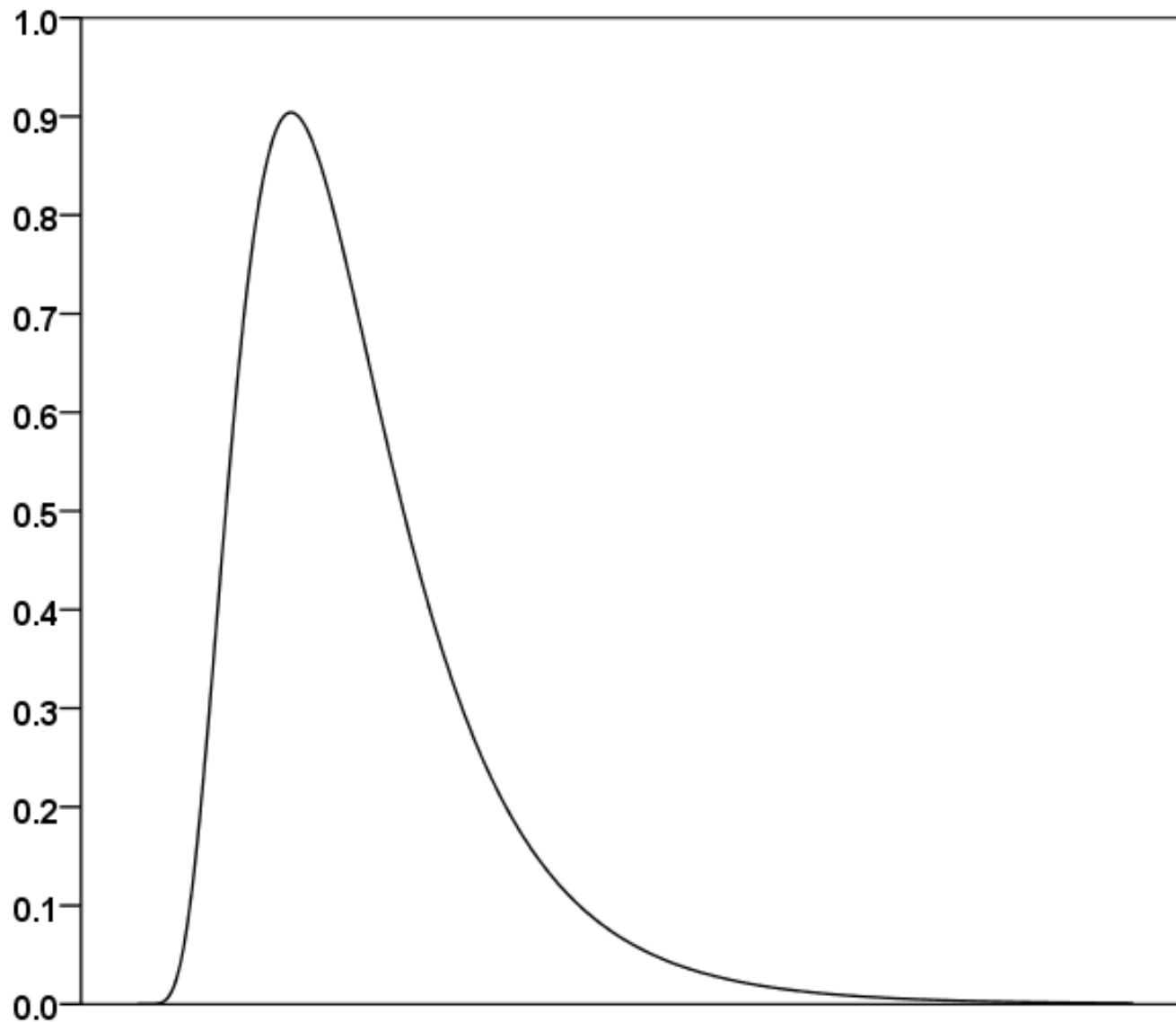
MEAN WIDTH OF INTERVALS

non-normal unbalanced data

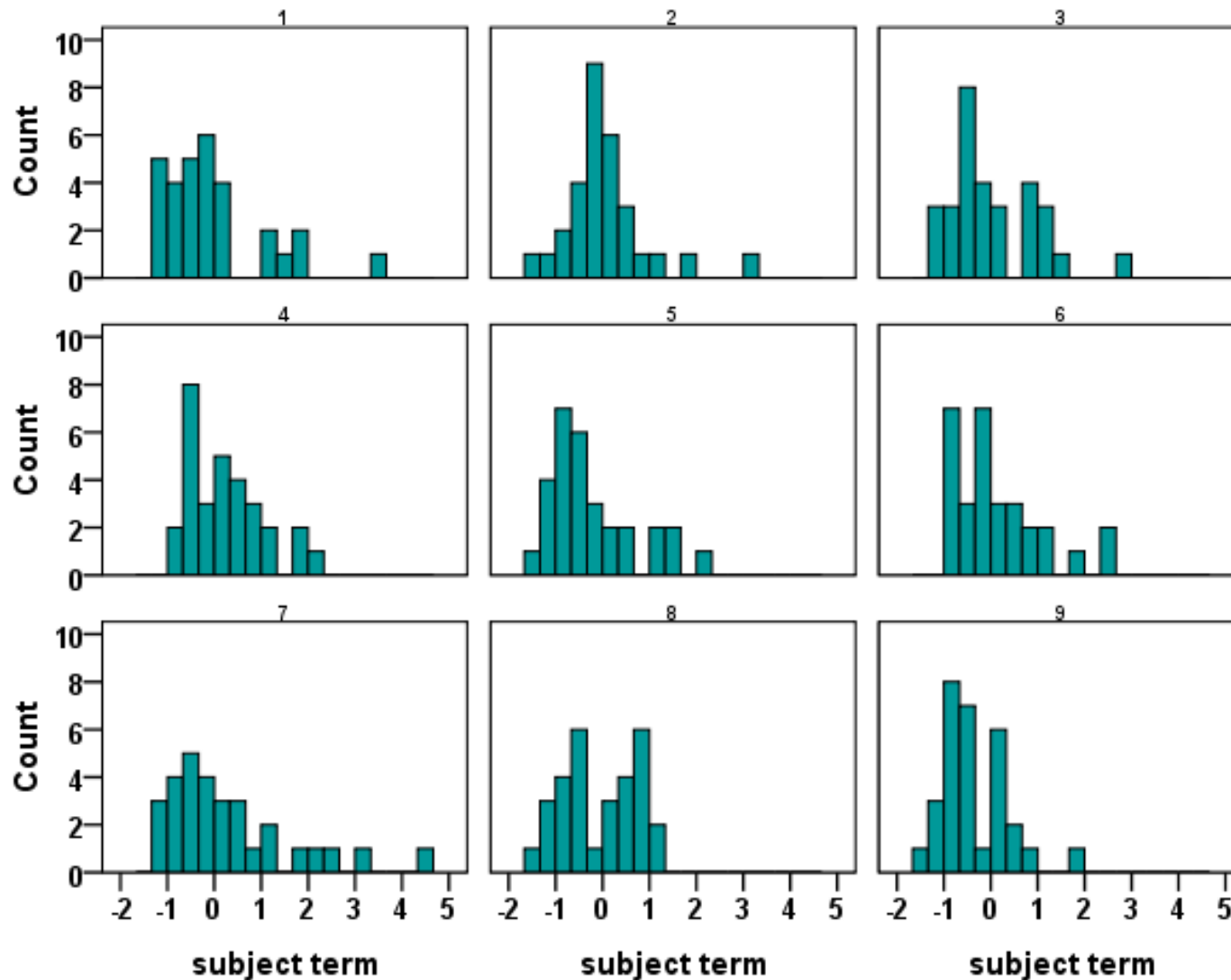


		Searle's		Bayesian
		mean width	mean +ve width	mean width
N=10 n=2	R=10	182.0	70.3	75.4
	R=90	66.1	57.5	52.0
N=10 n=5	R=10	69.3	50.2	54.8
	R=90	29.0	29.0	29.5
N=30 n=2	R=10	108.9	51.3	54.8
	R=90	23.7	23.7	23.9
N=10 n=5	R=10	40.7	31.7	33.1
	R=90	13.6	13.6	13.6

pdf of a lognormal(0,0.25)
skewness=1.75



Simulated non-normal subjects effects for the first 9 datasets, $N=30$, $n_i=2$, $R=10\%$





When is Searle's lower limit < 0 ?

$$\text{lower limit} = \frac{F/F_U - 1}{n + F/F_U - 1}$$

$$\text{-ve if } F = \frac{MS_{\text{subject}}}{MS_{\text{error}}} < F_U$$

For samples of $N=9, n=2$ $F_{N(n-1)=9}^{N-1=8} = 4.102$

For cadence $MS_{\text{subject}} = 442.9, MS_{\text{error}} = 134.0$

For 'reasonable' values of N & n , $2 < F_{N(n-1)}^{N-1} < 3$

N and $n \rightarrow \infty$ $F_{N(n-1)}^{N-1} \rightarrow 1$