

Web-based Supporting Materials for “Equivalence of binormal likelihood-ratio and bi-chi-squared ROC curve models” by Stephen L. Hillis

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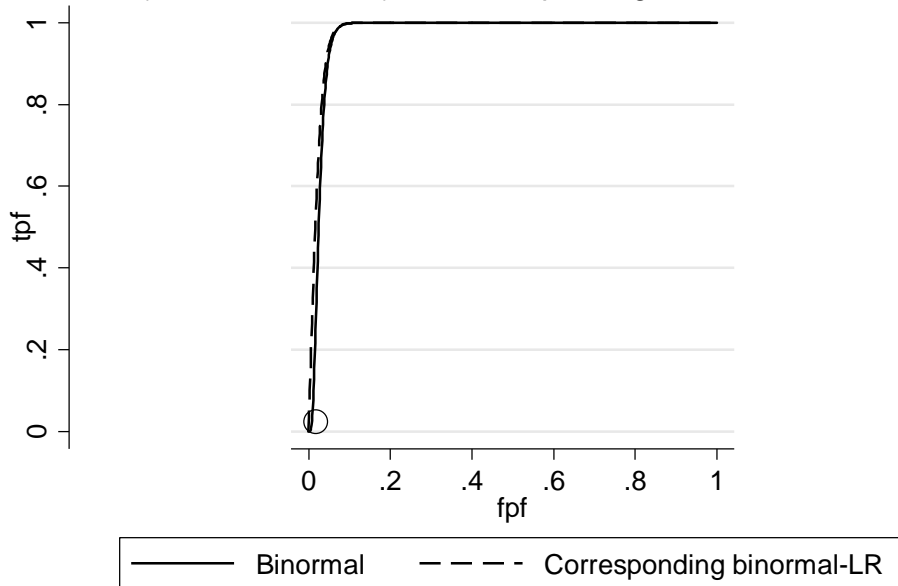
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Figure S1: Binormal ROC curve (with $b > 1$) and related plots

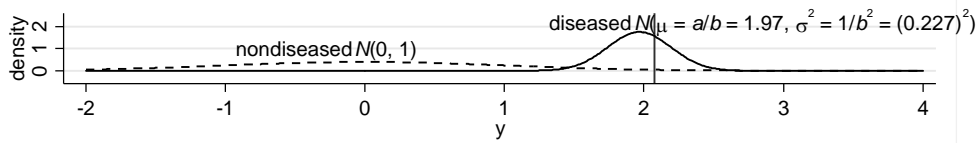
The binormal ROC curve (solid line) in (a) has an almost indiscernable hook in the lower left corner, as indicated by the circle. For the corresponding binormal random variable Y , the binormal conditional densities and the log-likelihood ratio are displayed in (b) and (c), respectively, where $c_1 = -ab/(1-b^2)$ is the threshold where the log-likelihood ratio attains its maximum. The bi-chi-squared conditional densities for $-Y^* = -(Y - c_1)^2$ are displayed in (d), and the binormal-LR ROC curve, based on $-Y^*$, is the dashed line in (a). Notes: the nondiseased and diseased densities are denoted by the dashed and solid lines, respectively, in (b) and (d); $\chi^2(1, \nu)$ denotes a chi-squared distribution with one degree of freedom and noncentrality parameter ν ; LR: likelihood ratio function; $\lambda = 1/b^2$; $\theta = a^2b^2/(1-b^2)^2$.

(NOTE: figure is on next page)

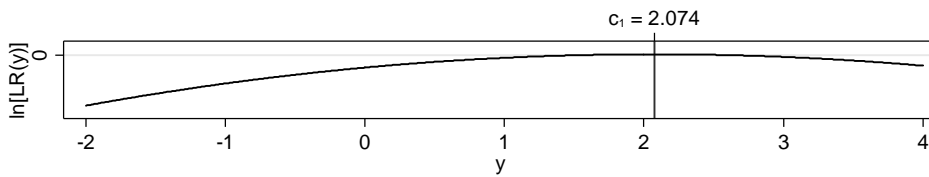
a) Binormal ($a = 8.65$, $b = 4.40$) and corresponding binormal-LR ROC curves



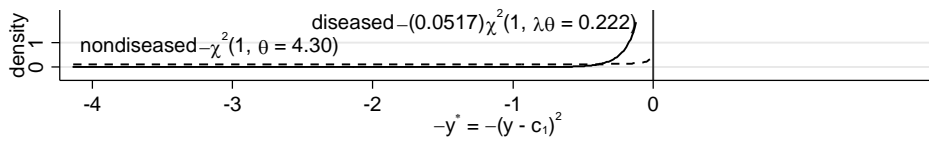
b) Binormal distribution for Y ($a = 8.65$, $b = 4.40$)



c) Log-likelihood ratio for Y



d) Bi-chi-squared distribution for $-Y^* = -(Y - c_1)^2$ ($\theta = 4.30$, $\lambda = 0.0517$)



Appendix S1: Simple bi-chi-squared program without macros

```

*****
This program is for the following paper: "Equivalence of binormal
likelihood-ratio and bi-chi-squared ROC curve models" by
Stephen L. Hillis.
*****
This program performs ML estimation for the bi-chi-square model and produces
the output for reader 1 using modality 1 in Table 4 for the Van Dyke et al data. This
program is written without macros to make it easier to understand.

Both of the AUC formulas in Table 3 are used and yield the same result (as they
should). The AUC formula from Table 3(d) is labeled "Hillis_AUC" and the formula
from Table 3(e) is labeled "Metz_AUC."
;
****VanDyke data -- these data can be downloaded from perception.radiology.uiowa.edu.
The data set name is VanDyke_anova.sas7bdat. Click on
perception.radiology.uiowa.edu --> Software --> Receiver Operating Characteristic (ROC)
--> SAS Programs to Perform MRMC Analysis. The file
can be found in "Installation Files for SAS MRMC Programs."
;
libname source "D:\Dorfman\data sets -- important";
**NOTE: specify above the directory where VanDyke data is stored**;

proc nlmixed data=source.VanDyke_anova gconv = 0 maxit = 500 ;
  where reader = 1 & treatment = 1;
  parms  delta1 .681 delta2 = .13  delta3 = .145 delta4 = .029
    log_lambda = 1.24 log_theta = 0.50 ; **deltas are the initial 4
rating-level nondiseased probabilities, e.g., delta1 = pr(R = 1|nondiseased);
  bounds  delta1-delta4 > 0;
  theta = exp(log_theta);
  lambda = exp(log_lambda);
  array delta[5] delta1 - delta5;
  array spec[5] spec1 - spec5;
  array fpf[5] fpf1 - fpf5;
  array tpf[5] tpf1 - tpf5;
  spec[1] = 0; **specificity = 0 for rating = 1**;
  temp = 0;
  do i = 2 to 5; **compute other specificities**;
    spec[i] = min(1,temp + delta[i-1]);
temp = spec[i];
  end;
  delta[5] = max(0, 1 - spec[5]);
  do i = 1 to 5;
    fpf[i] = 1 - spec[i];
  end;
  tpf[1] = 1;
  do i = 2 to 5;
    if fpf[i] = 0 or fpf[i] = 1 then do;
      if fpf[i] = 0 then tpf[i] = 0;
      if fpf[i] = 1 then tpf[i] = 1;
    end;
    else do;
      if lambda >= 1 then tpf[i] = 1 - probchi(lambda**i-1 * cinv(1 -
fpf[i],1,theta),1,lambda*theta);
      if lambda < 1 then tpf[i] = probchi(lambda**i-1 * cinv(fpf[i],1,theta),1,lambda*theta);
    end;
  end;
  if rating=5 then z= (truth = 0)*fpf[5] + (truth=1)*tpf[5];
  do i = 1 to 4;
    if rating = i then z= (truth = 0)*(fpf[i] -fpf[i+1]) + (truth=1)*(tpf[i] -tpf[i+1]);
  end;
  if z>1e-200 then ll = log(z); **to avoid undefined contribution**;

```

```

else ll= log(1e-200);
model rating ~ general(ll);
u1 = (lambda*sqrt(theta) - sqrt(theta))/sqrt(1+lambda);
u2 = (lambda*sqrt(theta) + sqrt(theta))/sqrt(1+lambda);
rho = (-1 + lambda)/(1 + lambda);
Metz_arg = sqrt(theta)*(-1+lambda)/sqrt(1+lambda); **this is d_a/sqrt(2)**;
*The next 3 statements avoid undefined values for the probbnrm and probnorm functions*;
if u1 < -4 then u1 = -4; if u1 > 4 then u1 = 4;
if u2 < -4 then u2 = -4; if u2 > 4 then u2 = 4;
if Metz_arg < -4 then metz_arg = -4; if metz_arg > 4 then metz_arg = 4;
**optional estimate statements below**;
*Note: AUC_Hillis and AUC_Metz use formulas (d) and (e) in Table 3, respectively*;
estimate "AUC_Hillis" (lambda>1)*(probbnrm(u1,u2,rho) + probbnrm(-u1,-u2,rho))
+ (lambda<1)*(1 - probbnrm(u1,u2,rho) - probbnrm(-u1,-u2,rho));
estimate "AUC_Metz" probnorm(metz_arg)
+ 2*probbnrm(-metz_arg,0,-2*sqrt(lambda)/(lambda+1));
estimate "lambda" lambda;
estimate "theta" theta;
estimate "a" sqrt((lambda - 1)**2 * theta) /sqrt(lambda);
estimate "b" 1/sqrt(lambda);
run;

```

Appendix S2: Bi-chi-squared program with macros

Bi-chi-squared program with macros for paper 10-05-14 revised 07-04-15b with Hillis and MetzAUC.sas;

*Appendix S2: Bi-chi-squared program with macros

This program is for the following paper: "The equivalence of the binormal likelihood-ratio and bi-chi-squared ROC curve models," by Stephen L. Hillis.

This program performs ML estimation for the bi-chi-square model and produces the output for Table 4 in the paper. Both of the AUC formulas in Table 3 are used and yield the same result (as they should). The AUC formula from Table 3(d) is labeled "Hillis_AUC" and the formula from Table 3(e) is labeled "Metz_AUC."

Output from the program is included.

The only statement that needs to be changed to run the program is the libname statment below -- the appropriate directory for the VanDyke data must be specified.

;

***VanDyke data -- these data can be downloaded from perception.radiology.uiowa.edu. The data set name is VanDyke_anova.sas7bdat. Click on perception.radiology.uiowa.edu --> Software --> Receiver Operating Characteristic (ROC) --> SAS Programs to Perform MRMC Analysis. The file can be found in "Installation Files for SAS MRMC Programs."

;

libname source "D:\Dorfman\data sets -- important"; *"D:\Dorfman\Obuchowski\Data sets";
IMPORTANT NOTE: specify above the directory where VanDyke data is stored;

%macro *BYCHI*;

```
proc nlmixed data=&dataset   gconv = 0 maxit = 500 ; where &where;
  ods output additionalEstimates = parmsout parameterEstimates = parm_estimates;
  %let numcat_minus_1 = %eval(&numcat - 1);
  parms  delta1 = &delta1 delta2 = &delta2  delta3 = &delta3 delta4 = &delta4
         log_lambda = &log_lambda log_theta = &log_theta ; **deltas are the initial p-1
rating level nondiseased probabilities, e.g., delta1 = pr(R = 1|nondiseaed);
  bounds delta1-delta&numcat_minus_1 > 0;
  theta = exp(log_theta);
  lambda = exp(log_lambda);
  array delta[&numcat] delta1 - delta&numcat;
  array spec[&numcat] spec1 - spec&numcat;
  array fpf[&numcat] fpf1 - fpf&numcat;
  array tpf[&numcat] tpf1 - tpf&numcat;
  spec[1] = 0; **specificity = 0 for rating = 1**;
  temp = 0;
  do i = 2 to &numcat; **compute other specificities**;
    spec[i] = min(1,temp + delta[i-1]);
  temp = spec[i];
  end;
  delta[&numcat] = max(0, 1 - spec[&numcat]);
  do i = 1 to &numcat;
    fpf[i] = 1 - spec[i];
  end;
  tpf[1] = 1;
  do i = 2 to &numcat;
    if fpf[i] = 0 or fpf[i] = 1 then do;
      if fpf[i] = 0 then tpf[i] = 0;
      if fpf[i] = 1 then tpf[i] = 1;
    end;
  end;
  else do;
```

```

        if lambda >= 1 then tpf[i] = 1 - probchi(lambda**(-1) * cinv(1 -
fpf[i],1,theta),1,lambda*theta);
        if lambda < 1 then tpf[i] = probchi(lambda**(-1) * cinv(fpf[i],1,theta),1,lambda*theta);
end;
end;
if rating=&numcat then z= (truth = 0)*fpf[&numcat] + (truth=1)*tpf[&numcat];
do i = 1 to &numcat_minus_1;
    if rating = i then z= (truth = 0)*(fpf[i] -fpf[i+1]) + (truth=1)*(tpf[i] -tpf[i+1]);
end;
if z>1e-200 then ll = log(z); **to avoid undefined contribution**;
    else ll= log(1e-200);
model rating ~ general(ll);
u1 = (lambda*sqrt(theta) - sqrt(theta))/sqrt(1+lambda);
u2 = (lambda*sqrt(theta) + sqrt(theta))/sqrt(1+lambda);
y =sqrt(theta);
rho = (-1 + lambda)/(1 + lambda);
Metz_arg = sqrt(theta)*(-1+lambda)/sqrt(1+lambda); **this is d_a/sqrt(2)**;
*The next 3 statements avoid undefined values for the probbnrm and probnorm functions*;
if u1 < -4 then u1 = -4; if u1 > 4 then u1 = 4;
if u2 < -4 then u2 = -4; if u2 > 4 then u2 = 4;
If Metz_arg <-4 then metz_arg = -4; if metz_arg > 4 then metz_arg = 4;
**optional estimate statements below**;
*Note: AUC_Hillis and AUC_Metz use formulas (d) and (e) in Table 3, respectively.*;
estimate "AUC_Hillis" (lambda>1)*(probbnrm(u1,u2,rho) + probbnrm(-u1,-u2,rho))
    + (lambda<1)*(1 - probbnrm(u1,u2,rho) - probbnrm(-u1,-u2,rho));
estimate "AUC_Metz" probnorm(metz_arg)
    + 2*probbnrm(-metz_arg,0,-2*sqrt(lambda)/(lambda+1));
estimate "lambda" lambda;
estimate "theta" theta;
estimate "a" sqrt((lambda - 1)**2 * theta) /sqrt(lambda);
estimate "b" 1/sqrt(lambda);

run;
data parmsout&reader.&treatment; set parmsout; reader = &reader; treatment = &treatment; run;
%mend BYCHI;

data VanDyke; set source.VanDyke_anova; run;

****This next part is only for determining the empirical fpf values for ratings 1-4 that will be
used as the initial delta values in the program*****;

ods listing close; **this statement suppresses the printing of output
not needed for the bi-chi-square program**;
proc freq data = vandyke ;
ods output crosstabfreqs = freqtable;
table treatment*reader*truth*rating; run;
proc print data = freqtable; run;
data freqtable1; set freqtable; if truth = 0; if rating ne .;
keep treatment reader rating rowpercent;
data freqtable2; set freqtable1;
delta = rowpercent/100;
if rating ne 5;
ods listing; **now printing turned back on**;
proc print; format delta 5.3;
title "Starting values for delta";
title3 "Zero values will be changed to .001";
title4 "and totals constrained to not exceed 1";
title5 "(by subtracting a little from large deltas)";
TITTLE6 "as indicated in bi-chi-square program macros statements below";
run;
title "";

/* below are results of above print statement

```

Starting values for delta

Zero values will be changed to .001
and totals constrained to not exceed 1
(by subtracting a little from large deltas)
as indicated in bi-chi-square program macro statements below

Table with 6 columns: Obs, treatment, reader, rating, Row Percent, delta. It lists 40 observations with their respective treatment, reader, rating, and calculated delta values.

*****end of empirical fpf value determination section*****;
*/

*****starting estimates from binormal fit*****

Below are the estimated binormal a and b values and AUC for the VanDyke data and the corresponding lambda, theta, log(lambda) and log(theta) values. The log(lambda) and log(theta) values are used as the initial starting values in the bi-chi-square program.

These binormal estimates were obtained using the software OR/DBM MRMC for SAS 3.0 which is available for download from perception.radiology.uiowa.edu.

*****;

/*

Obs	treatment	reader	AUC	a	b	lambda	theta	log_ lambda	log_ theta
1	1	1	0.93316	1.7022	0.5368	3.4707	1.6473	1.24435	0.49917
2	1	2	0.88957	1.4037	0.5610	3.1778	1.3202	1.15618	0.27782
3	1	3	0.92920	1.7409	0.6346	2.4830	3.4216	0.90945	1.23012
4	1	4	0.97046	1.9255	0.2015	24.6279	0.1636	3.20388	-1.81060
5	1	5	0.83259	1.0630	0.4635	4.6545	0.3938	1.53784	-0.93188
6	2	1	0.95082	1.8502	0.5030	3.9525	1.5520	1.37436	0.43956
7	2	2	0.93460	1.6552	0.4473	4.9976	0.8568	1.60897	-0.15461
8	2	3	0.92755	1.6220	0.4878	4.2017	1.0783	1.43549	0.07542
9	2	4	1.00000	7.1233	0.8806	1.2896	780.1023	0.25435	6.65943
10	2	5	0.94481	1.7329	0.4221	5.6124	0.7922	1.72499	-0.23298

*/

Below are the statements for calling the bi-chi-square algorithm macro to obtain the bi-chi-square results given in the paper.

*****;

```
%let numcat = 5;

%let dataset = vandyke;
%let log_lambda = 1.24; %let log_theta = 0.50;
%let reader = 1; %let treatment = 1;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .681;
%let delta2 = .13;
%let delta3 = .145;
%let delta4 = .029;
%BYCHI;

%let log_lambda = 1.16; %let log_theta = 0.28;
%let reader = 2; %let treatment = 1;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .001;
%let delta2 = .87;
%let delta3 = .087;
%let delta4 = .029;
%BYCHI;

%let log_lambda = .91; %let log_theta = 1.23;
%let reader = 3; %let treatment = 1;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .304;
%let delta2 = .507;
%let delta3 = .072;
%let delta4 = .087;
%BYCHI;

%let log_lambda = 3.20; %let log_theta = -1.81;
%let reader = 4; %let treatment = 1;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .89; *changed from .899*;
%let delta2 = .043;
%let delta3 = .058;
%let delta4 = .001; *changed from 0*;
%BYCHI;

%let log_lambda = 1.54; %let log_theta = -0.93;
%let reader = 5; %let treatment = 1;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .565;
%let delta2 = .275;
```

```

%let delta3 = .13;
%let delta4 = .014;
%BYCHI;

%let log_lambda = 1.37; %let log_theta = 0.44;
%let reader = 1; %let treatment = 2;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .333;
%let delta2 = .348;
%let delta3 = .217;
%let delta4 = .101;
%BYCHI;

%let log_lambda = 1.61; %let log_theta = -0.15;
%let reader = 2; %let treatment = 2;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .087;
%let delta2 = .812;
%let delta3 = .099; *changed from 0.101;
%let delta4 = .001; *changed from 0.000*;
%BYCHI;

%let log_lambda = 1.44; %let log_theta = 0.08;
%let reader = 3; %let treatment = 2;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .362;
%let delta2 = .449;
%let delta3 = .116;
%let delta4 = .058;
%BYCHI;

%let log_lambda = 0.25; %let log_theta = 6.66;
%let reader = 4; %let treatment = 2;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .638;
%let delta2 = .304;
%let delta3 = .055; *changed from 0.58*;
%let delta4 = .001; *changed from 0.000*;
%BYCHI;

%let log_lambda = 1.72; %let log_theta = -0.23;
%let reader = 5; %let treatment = 2;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .304;
%let delta2 = .565;
%let delta3 = .128; *changed from 0.130*;
%let delta4 = .001; *changed from 0.000*;
%BYCHI;

data parmsout_all; set parmsout11 parmsout21 parmsout31 parmsout41 parmsout51
  parmsout12 parmsout22 parmsout32 parmsout42 parmsout52;
  keep label estimate reader treatment; run;
proc transpose data = parmsout_all out = parmsout_all_tr; id label; by treatment reader;
  run;

options ls = 80;
proc print data = parmsout_all_tr; format AUC_Hillis AUC_Metz a b lambda theta 15.6;
  id treatment reader; var auc_hillis auc_metz lambda theta a b ;run;

```

Below are the results from the above print statement, which include the Bi-chi-square AUC, lambda, and theta values for Table 4.

treatment	reader	AUC_Hillis	AUC_Metz	lambda	theta
1	1	0.934040	0.934040	3.418921	1.706011
1	2	0.891071	0.891071	3.172871	1.324855
1	3	0.928886	0.928886	2.532211	3.239212
1	4	0.977460	0.977460	786.713272	0.000017
1	5	0.840559	0.840559	9.366031	0.059426
2	1	0.951936	0.951936	3.788976	1.697362
2	2	0.925992	0.925992	73.215915	0.000022
2	3	0.930432	0.930432	3.940212	1.234458
2	4	0.999937	0.999968	1.283937	780.544368
2	5	0.942687	0.942687	12.075803	0.217395

treatment	reader	a	b
1	1	1.708709	0.540823
1	2	1.404080	0.561402
1	3	1.732959	0.628420
1	4	0.114029	0.035653
1	5	0.666394	0.326755
2	1	1.866685	0.513735
2	2	0.039241	0.116868
2	3	1.645724	0.503779
2	4	7.000815	0.882527
2	5	1.486081	0.287768

Below are results when I use different starting values for reader 3 and modality 1, which gives me the same answer as PROPROC;

```

%let numcat = 5;
%let dataset = vandyke;
%let log_lambda = -4 -2 0 2 4 ;
%let log_theta = -4 -2 0 2 4 ;
%let reader = 3; %let treatment = 1;
%let where = reader = &reader & treatment = &treatment;
%let delta1 = .304;
%let delta2 = .507;
%let delta3 = .072;
%let delta4 = .087;
%BYCHI;

data parmsout31; set parmsout31; keep label estimate reader treatment; run;
proc transpose data = parmsout31 out = parmsout31_tr; id label; by treatment reader;
run;
proc print data = parmsout31_tr; format AUC_hillis AUC_Metz a b lambda theta 15.6;
id treatment; var auc_hillis auc_metz lambda theta a b ;run;

```

**Results from above print statement:

treatment	AUC_Hillis	AUC_Metz	lambda
1	0.907833	0.907833	46.925054

treatment	theta	a	b
1	0.000056	0.049949	0.145981

;