# **Examples for Testing of the Indirect Estimation Prototype**

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#### Introduction

This document contains a set of examples that demonstrates the estimation capabilities of the Indirect Estimation Prototype. These capabilities are demonstrated by a set of examples that are presented in increasing level of complexity.

Each example consists of a visual description of the model to be estimated. The states are numbered and transitions are noted by coefficient names. In case the model uses a coefficient table depending on a covariate, the table is presented after the graphic model. The coefficients are the unknowns to be deduced during estimation.

To estimate the model, each example provides a set of studies that provide sufficient information to estimate the model. Each study is presented graphically in model terminology, i.e. the start and end state of the study is provided using the states declared in the model. The information for each study consists of:

- T Study length in time units
- Study related information of one of the following forms:
  - o Studies that provide data
    - $\blacksquare$  N Initial population at start state
    - x Incident count reaching the study end state by the end of study
  - o Studies that provide regression information
    - $F(t,\lambda)$  Regression formula
    - $\hat{\lambda}$  Regression coefficients provided by the study
    - $\Sigma$  The covariance matrix associated with these coefficients
- Population In some cases, population information is required to interpret the study or to support the system operation. The population is provided per study and may be presented in one of the following ways:
  - o Default theoretically not required, although the software may require some definition for proper functionality.
  - o Dummy Any population that describes one or more covariates to allow proper functionality of the software. Not required theoretically.
  - o Distribution a set of covariates and their marginal distribution function
  - Data a table that its columns are covariates and each row contains the values for this column for each individual.

The expected outcomes that describe the optimal coefficient values are presented at the end of each example. In some cases, some comments or variations on the example are also provided as well as the expected partial likelihood related to a specific study.

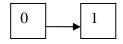
For further information about the project, please visit the project web site: <a href="http://www.med.umich.edu/mdrtc/cores/DiseaseModel/">http://www.med.umich.edu/mdrtc/cores/DiseaseModel/</a>

# **Example 1:** Simple Example with Trivial Outcome:

## Model

$$0 P_0$$
 1

# Study 1



Data: N=100 x=20 T=3

Population: Default - Theoretically none needed

Partial Likelihood:

$$[1-(1-p01)^{\Lambda}T]^{\Lambda}x\ [(1-p01)^{\Lambda}T]^{\Lambda}(N-x)\ [1-(1-p01)^{\Lambda}3]^{\Lambda}20\ [(1-p01)^{\Lambda}3]^{\Lambda}(80)$$

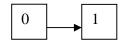
$$\hat{p}_0 = 1 - (1 - \frac{x}{N})^{1/t} = 0.0717$$

# **Example 2:** Theoretical Example with previous Algorithms (based on Isaman, 2006, Table 1, model 1:1:3)

#### Model

$$0$$
  $P_0$   $1$   $P_1$   $2$ 

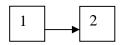
#### Study 1



Data: N=500 x=204.755 T=5 **Population:** Default - Theoretically none needed

**Partial Likelihood**:  $[1-(1-P0)^T]^x [(1-P0)^T]^(N-x)$ 

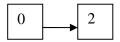
#### Study 2



Data: N=500 x=336.16 T=5 **Population:** Default - Theoretically none needed

**Partial Likelihood**:  $[1-(1-P1)^T]^x [(1-P1)^T]^(N-x)$ 

### **Study 3-5**



Data Study 3: N=500 x=73.35 T=5

Population: Default - Theoretically none needed

**Partial Likelihood**:  $[1-(1-P)]^x [(1-P)]^n(N-x)$  where

 $P = P0*P1+P0*P1*(1-P0)+P0*P1*(1-P1)+P0*P1*(1-P0)^2+P0*P1*(1-P1)^2+P0*P1*(1-P0)^2+P0*P1*(1-P1)+P0*P1*(1-P0)^3+P0*P1*(1-P1)^3+P0*P1*(1-P0)^2+P0*P1*(1-P1)^2+P0*P1*(1-P0)^2*(1-P1)$ 

#### **Expected Outcome:**

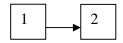
 $\hat{p}_0 = 0.1$   $\hat{p}_1 = 0.2$ 

# **Example 3:** Theoretical Example - Missing Some Data- (based on Isaman, 2006, Table 1, model 0:1:3)

### Model

$$0$$
  $P_0$   $1$   $P_1$   $2$ 

# Study 1

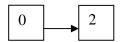


Data: N=500 x=336.16 T=5

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 2-4



Data Study 3: N=500 x=73.35 T=5 Data Study 4: N=400 x=58.68 T=5 Data Study 5: N=300 x=44.01 T=5

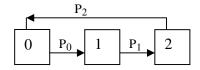
Population: Default - Theoretically none needed

Partial Likelihood: Not given

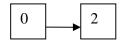
$$\hat{p}_0 = 0.1$$
  $\hat{p}_1 = 0.2$ 

# **Example 4:** Funny Loop Example – No Direct Data – (from Isaman, 2006, Table 2, model C)

#### Model



#### **Study 1-3**



**Population:** Default - Theoretically none needed

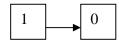
**Partial Likelihood**:  $[1-(1-P)]^x [(1-P)]^(N-x)$  where

$$\begin{split} P &= P0*P1 + P0*P1*(1-P0) + P0*P1*(1-P1) + P0*P1*(1-P0)^2 + P0*P1*(1-P1)^2 + P0*P1*(1-P0)^2 + P0*P1*(1-P$$

P0)^3+P0\*P1\*(1-P1)^3+P0\*P1\*(1-P0)\*(1-

P1)^2+P0\*P1\*(1-P0)^2\*(1-P1)

#### **Study 4-6**



**Population:** Default - Theoretically none needed

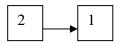
**Partial Likelihood**:  $[1-(1-P)]^x [(1-P)]^(N-x)$  where

 $P = P2*P1+P2*P1*(1-P2)+P2*P1*(1-P1)+P2*P1*(1-P2)^2+P2*P1*(1-P1)^2+P2*P1*(1-P2)^$ 

P2)^3+P2\*P1\*(1-P1)^3+P2\*P1\*(1-P2)\*(1-

P1)^2+P2\*P1\*(1-P2)^2\*(1-P1)

# **Study 7-9**



Data Study 7: N=500 x=99.15 T=5
Data Study 8: N=400 x=79.32 T=5
Data Study 9: N=300 x=59.49 T=5

Population: Default - Theoretically none needed

**Partial Likelihood**:  $[1-(1-P)]^{x} [(1-P)]^{(N-x)}$  where

$$\begin{split} P &= P0*P2 + P0*P2*(1-P0) + P0*P2*(1-P2) + P0*P2*(1-P0)^2 + P0*P2*(1-P2)^2 + P0*P2*(1-P0)^2 + P0*P2*(1-P$$

P0)^3+P0\*P2\*(1-P2)^3+P0\*P2\*(1-P0)\*(1-

P2)^2+P0\*P2\*(1-P0)^2\*(1-P2)

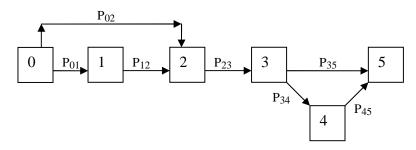
$$\hat{p}_0 = 0.1$$

$$\hat{p}_1 = 0.2$$

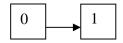
$$\hat{p}_2 = 0.3$$

# **Example 5:** Example with a Pooled State

#### Model



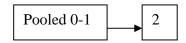
### **Study 1-2**



Population: Default - Theoretically none needed

Partial Likelihood: Not given

### Study 3

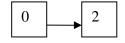


Data: N=398 x=79.18810184 T=4, Prevalence = 50% State 0

Population: Default - Theoretically none needed

Partial Likelihood: Not given

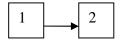
#### Study 4



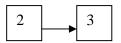
Data: N=176 x=16.118751520256 T=6 **Population:** Default - Theoretically none needed

Partial Likelihood: Not given

### **Study 5-6**

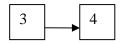


Population: Default - Theoretically none needed



Partial Likelihood: Not given

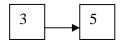
#### Study 8



Data: N=1000 x=40 T=1 **Population:** Default - Theoretically none needed

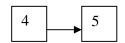
Partial Likelihood: Not given

# **Study 9-10**



Partial Likelihood: Not given

### Study 11



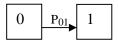
Data: N=23 x=5.2030384375 T=5 **Population:** Default - Theoretically none needed

Partial Likelihood: Not given

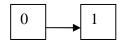
$$\hat{p}_{01} = 0.03$$
,  $\hat{p}_{02} = 0.01$ ,  $\hat{p}_{12} = 0.1$ ,  $\hat{p}_{23} = 0.02$ ,  $\hat{p}_{34} = 0.04$ ,  $\hat{p}_{35} = 0.2$ ,  $\hat{p}_{45} = 0.05$ 

# **Example 6:** Simple Example with Covariates:

## Model



## Study 1



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	90	100
T= 3	30	20

**Population:** Any population defining Gender in the range [0,1]

**Partial Likelihood**:  $[1-(1-P01)^T]^x$ male \*  $[(1-P01)^T]^x$ (Nmale-xmale)

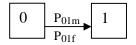
 $[1-(1-P01)^T]^x$ female\* $[(1-P01)^T]^(N$ female-xfemale)

=[1-(1-P01)^3]^20 [(1-P01)^3]^80 [1-(1-P01)^3]^30 [(1-P01)^3]^60

$$\hat{p}_{01} = 1 - (1 - \frac{x_{male} + x_{female}}{N_{male} + N_{female}})^{1/t} = 0.09678$$

## **Example 7:** Non Unique Solution - Example with Covariates #2:

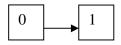
#### Model



Model coefficient table for the transition from 0 to 1:

Gender = 0	Gender = 1
P01f	P01m

#### Study 1

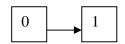


Data: N=100 = 75.2250 = T=3**Population:** Gender ~ Bernoulli(0.75)

**Partial Likelihood**:  $[Wm*(1-(1-P01m)^T) + (1-Wm)*(1-(1-P01f)^T)]^x + (1-Wm)^2 + (1-Wm$ 

[Wm\*(  $(1-P01m)^T$ ) +  $(1-Wm)^*$ (  $(1-P01f)^T$ )]^(N-x) Where Wm is the prevalence of male, i.e. Wm = 0.75

## Study 2



Data: N=100 = 54.2500 = T=2**Population:** Gender ~ Bernoulli(0.25)

**Partial Likelihood**:  $[Wm*(1-(1-P01m)^T) + (1-Wm)*(1-(1-P01f)^T)]^x +$ 

[Wm\*(  $(1-P01m)^T$ ) +  $(1-Wm)^*$ (  $(1-P01f)^T$ )]^(N-x) Where Wm is the prevalence of male, i.e. Wm = 0.25

#### **Expected Outcome:**

$$\hat{p}_{01f} = 0.3$$

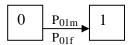
$$\hat{p}_{01m} = 0.4$$

#### Simple variation to demonstrate the estimably:

If the same example is run without study 2, then the result is inestimable since the model requires more information than the study supplies. The likelihood graph will show a ridge, rather than a peak, meaning that there are multiple solutions. The system will return one such solution that may vary depending on the initial guess.

## **Example 8:** Simple Example with Covariates #3:

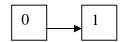
#### Model



Model coefficient table for the transition from 0 to 1:

Gender = 0	Gender = 1
P01f	P01m

### Study 1



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	90	100
T=3	30	20

**Population:** Any population that includes Gender in the range [0,1]

**Partial Likelihood**: [1-(1-P01m)^T]^xmale \* [(1-P01m)^T]^(Nmale-xmale)

 $[1-(1-P01f)^T]^x$ female\* $[(1-P01f)^T]^(N$ female-xfemale)

=[1-(1-P01m)^3]^20 [(1-P01m)^3]^80 [1-(1-P01f)^3]^30 [(1-P01f)^3]^60

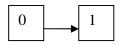
$$\hat{p}_{01m} = 1 - (1 - \frac{x}{N})^{1/t} = 0.07168$$

$$\hat{p}_{01f} = 1 - (1 - \frac{x}{N})^{1/t} = 0.12642$$

# **Example 9:** Simple Example with Regression Outcome:

### Model

### Study 1



Data:  $F(t, \lambda) = e^{-\lambda t}$ ,  $\hat{\lambda} = 0.1$ ,  $\sigma^2 = 0.05$ , T=5

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# **Expected Outcome:**

$$\hat{p}_{01} = 1 - (1 - e^{-0.1*5})^{1/5} = 0.17018$$

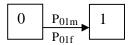
# Simple variation for authentication:

For the simple example where T=1 the result should be:

$$\hat{p}_{01} = e^{-0.1} = 0.90484$$

## **Example 10:** Simple Example with Regression Outcome and Covariates:

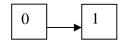
#### Model



Model coefficient table for the transition from 0 to 1:

Gender = 0	Gender = 1
P01f	P01m

#### Study 1



Data:

$$F(t,\lambda) = e^{-(\lambda_0 + \lambda_1 Male + \lambda_0 BP)t},$$

$$F(t,\lambda) = e^{-(\lambda_0 + \lambda_1 Male + \lambda_0 BP)t},$$

$$\hat{\lambda} = (0.1,0.2,0.05), \Sigma = \begin{pmatrix} 1 & 0 & 0 \\ 0 & 1 & 0 \\ 0 & 0 & 1 \end{pmatrix}, T=1,2,3$$

**Population:** 4 individuals (Gender, BP) = (0,125), (0,115), (1,125), (1,115)Where Gender =1 means male

Partial Likelihood: Not given

#### **Expected Outcome:**

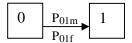
$$\begin{split} \hat{p}_{01m} &= 1 - \left(1 - (1/n_{Male}) \sum_{i \ni Gender_i = Male = 1} e^{\left(-\hat{\lambda}_0 - \hat{\lambda}_1 * Gender_i - \hat{\lambda}_2 * BP_i\right)T} \right)^{1/T} \approx \\ 1 - (1 - e^{\left(-\hat{\lambda}_0 - \hat{\lambda}_1 * Gender - \hat{\lambda}_2 * average(BP|Gender = Male)\right)T})^{1/T} = 1 - (1 - e^{\left(-0.1 - 0.2 - 0.05 * 120\right)T})^{1/T} = 1 - (1 - e^{-6.3T})^{1/T} \\ \hat{p}_{01f} &= 1 - \left(1 - (1/n_{Female}) \sum_{i \ni Gender_i = Female = 0} e^{\left(-\hat{\lambda}_0 - \hat{\lambda}_1 * Gender_i - \hat{\lambda}_2 * BP_i\right)T}\right)^{1/T} \approx \\ 1 - (1 - e^{\left(-\hat{\lambda}_0 - \hat{\lambda}_1 * Gender - \hat{\lambda}_2 * average(BP|Gender = Female)\right)T})^{1/T} = 1 - (1 - e^{\left(-0.1 - 0.05 * 120\right)T})^{1/T} = 1 - (1 - e^{-6.1T})^{1/T} \end{split}$$

T	P <sub>01 m</sub>	P <sub>01 f</sub>
1	0.0018939888023991047	0.0023133231471746951
2	1.9011877737673544e-06	2.8362402018089483e-06
3	2.672246868229422e-09	4.8691513043763734e-09
1 Approx.	0.00183630477703	0.00224286771949
2 Approx.	1.6860090383818971e-06	2.5152309667264561e-06
3 Approx.	2.0640158471252334e-09	3.7608821612522547e-09

**Note**: The approximation gives an idea for results with similar populations

## **Example 11:** Simple Example with Regression Outcome Consistent with Time:

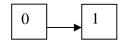
#### Model



Model coefficient table for the transition from 0 to 1:

Gender = 0	Gender = 1
P01f	P01m

#### Study 1



Data:

$$F(t,\lambda) = 1 - e^{-(\lambda_0 + \lambda_1 Male + \lambda_0 BP)t}$$

$$F(t,\lambda) = 1 - e^{-(\lambda_0 + \lambda_1 Male + \lambda_0 BP)t},$$

$$\hat{\lambda} = (.1,.2,.005), \Sigma = \begin{pmatrix} 1 & 0 & 0 \\ 0 & 1 & 0 \\ 0 & 0 & 1 \end{pmatrix}, T=1,2,3$$

4 individuals (Gender, BP) = (0,125), (0,115), (1,125), (1,115)

Where Gender = 1 means male

Partial Likelihood: Not given

#### **Expected Outcome:**

$$\begin{split} \hat{p}_{01m} &= 1 - \left( (1/n_{Male}) \sum_{i \ni Gender_i = Male = 1} e^{\left( -\hat{\lambda}_0 - \hat{\lambda}_1^* * Gender_i - \hat{\lambda}_2^* * BP_i \right) T} \right)^{1/T} \approx \\ 1 - e^{\left( -\hat{\lambda}_0 - \hat{\lambda}_1^* * Gender - \hat{\lambda}_2^* * average(BP|Gender = Male) \right)} = 1 - e^{\left( -0.1 - 0.2 - 0.005*120 \right)} = 1 - e^{\left( -0.9 \right)} \\ \hat{p}_{01f} &= 1 - \left( (1/n_{Female}) \sum_{i \ni Gender_i = Female = 0} e^{\left( -\hat{\lambda}_0 - \hat{\lambda}_1^* * Gender_i - \hat{\lambda}_2^* * BP_i \right) T} \right)^{1/T} \approx \\ 1 - e^{\left( -\hat{\lambda}_0 - \hat{\lambda}_1^* * Gender - \hat{\lambda}_2^* * average(BP|Gender = Female) \right)} = 1 - e^{\left( -0.1 - 0.005*120 \right)} = 1 - e^{\left( -0.7 - 0.005*120 \right)} = 1 - e^{\left( -0.7 - 0.005*120 \right)} \end{split}$$

T	P <sub>01 m</sub>	P <sub>01 f</sub>
1	0.59330328062324944	0.50325950521854423
2	0.59317626067050078	0.5031043626979157
3	0.59304935961526495	0.50294936539903712
Approx.	0.59343034025940089	0.50341469620859047

**Note**: The approximation gives an idea for results with similar populations

The Jacobian and the Covariance matrices for T=1 should be:

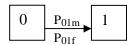
$$J = \begin{bmatrix} 0.40669672 & 0.40669672 & 48.75277982 \\ 0.49674049 & 0 & 59.54677974 \end{bmatrix}$$

$$\Sigma_{New} = J\Sigma J^{T} = \begin{bmatrix} 2377.16434498 & 2903.27306483 \\ 2903.27306483 & 3546.06572908 \end{bmatrix}$$

Note that row order in the Jacobian and diagonal elements in the covariance matrix may switch if the output vector is ordered by the program [female, male] rather than [male, female].

## **Example 12:** Regression with UKPDS formula:

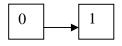
#### Model



Model coefficient table for the transition from 0 to 1:

Gender = 0	Gender = 1
P01f	P01m

#### Study 1



$$\text{Data:} \ \ F(\lambda) = 1 - e^{-\lambda_1 \lambda_2^{(AGE-55)} \lambda_3^{(1-MALE)} \lambda_4^{(RACE)} \lambda_5^{(SMOKE)} \lambda_6^{(BP-135.7)/10} \left( \frac{1 - \lambda_7^T}{1 - \lambda_7} \right) }$$

 $\hat{\lambda} = \begin{bmatrix} 0.0112 & 1.0590 & 0.5250 & 0.3900 & 1.3500 & 1.0880 & 1.078 \end{bmatrix}$ 

 $\Sigma = Diag \left( 2.0408*10^{-6} \quad 3.1497*10^{-5} \quad 0.0028699 \quad 0.010412 \quad 0.014994 \quad 0.00070387 \quad 0.00026656 \right)$ 

**Population:** 4 individuals (Age, Gender, Race, Smoke, BP)

54,1,0,0,115

56,1,1,0,125

54,0,0,1,115

56,0,0,0,125

Where Gender =1 means male

### Partial Likelihood: Not given

## **Expected Outcome:**

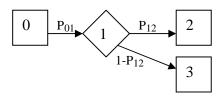
$$\begin{split} \hat{p}_{01m} &= 1 - \left( (1/n_{Male}) \sum_{i \ni Gender_i = Male = 1}^{-\lambda_1 \lambda_2^{(AGE - 55)} \lambda_3^{(1-MALE)} \lambda_4^{(RACE)} \lambda_5^{(SMOKE)} \lambda_6^{(BP - 135.7)/10} \left( \frac{1 - \lambda_7^T}{1 - \lambda_7} \right) \right)^{1/T} \\ \hat{p}_{01f} &= 1 - \left( (1/n_{Female}) \sum_{i \ni Gender_i = Female = 0}^{-\lambda_1 \lambda_2^{(AGE - 55)} \lambda_3^{(1-MALE)} \lambda_4^{(RACE)} \lambda_5^{(SMOKE)} \lambda_6^{(BP - 135.7)/10} \left( \frac{1 - \lambda_7^T}{1 - \lambda_7} \right) \right)^{1/T} \end{split}$$

T	P <sub>01 m</sub>	P <sub>01 f</sub>
1	0.00653004287191838	0.00597431015512395
2	0.00678083530428653	0.00620653272582317
3	0.007044275112843	0.00645076680543022

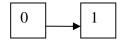
Note that for time = 1 the formula is simplified.

# **Example 13:** Event States – Basic Test

### Model

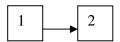


# Study 1



Partial Likelihood: Not given

# Study 2



Data: N=1000 x=300 T=1

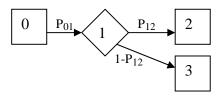
Population: Default - Theoretically none needed

Partial Likelihood: Not given

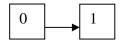
$$\hat{p}_{01} = 0.2$$
  $\hat{p}_{12} = 0.3$ 

# **Example 14:** Event States – Event Bypass Study

### Model



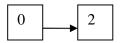
# Study 1



Data: N=1000 x=200 T=1 **Population:** Default - Theoretically none needed

Partial Likelihood: Not given

# Study 2



Data: N=1000 x=60 T=1

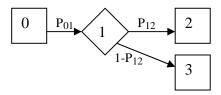
Population: Default - Theoretically none needed

Partial Likelihood: Not given

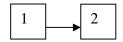
$$\hat{p}_{01} = 0.2$$
  $\hat{p}_{12} = 0.3$ 

# **Example 15:** Event States – Another Event Bypass

#### Model



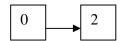
# Study 1



Data: N=1000 x=300 T=1 **Population:** Default - Theoretically none needed

Partial Likelihood: Not given

# Study 2



Data: N=1000 x=60 T=1 **Population:** Default - Theoretically none needed

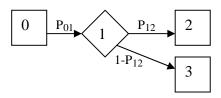
Partial Likelihood: Not given

$$\hat{p}_{01} = 0.2$$

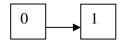
$$\hat{p}_{12} = 0.3$$

# **Example 16:** Event States – Two year study

#### Model



# Study 1

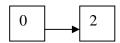


Data: N=1000 x=360=(200+160) T=2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 2



Data: N=1000 x=108=(60+48) T=2

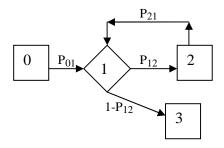
Population: Default - Theoretically none needed

Partial Likelihood: Not given

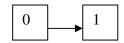
$$\hat{p}_{01} = 0.2$$
  $\hat{p}_{12} = 0.3$ 

# **Example 17:** Event States – Loop Back To Event

#### Model



## Study 1

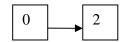


Data: N=1000 x=360 T=2x=1000\*0.2+1000\*(1-0.2)\*0.2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

## Study 2

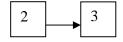


Data: N=10000 x=1080 T=2x=10000\*0.2\*0.3 +10000\*(1-0.2)\*0.2\*0.3

Population: Default - Theoretically none needed

Partial Likelihood: Not given

#### Study 3



Data:  $N=10000 \quad x=4816 \quad T=2 \\ x=10000*(0.4*0.7) +10000*(1-0.4*0.7)*(0.4*0.7)$ 

Population: Default - Theoretically none needed

Partial Likelihood: Not given

#### **Expected Outcome:**

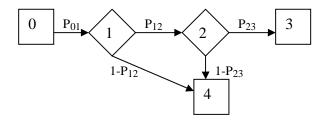
$$\hat{p}_{01} = 0.2$$
  $\hat{p}_{12} = 0.3$   $\hat{p}_{21} = 0.4$ 

## Simple variation for authentication:

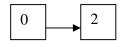
The same results should be reported for the same studies where T=1 and the study outcomes for studies 1-3 are 200, 600, 2800 respectively.

## **Example 18:** Event States – Multiple Event States

#### Model



### Study 1

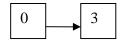


Data: N=1000 x=108 T=2 x=1000\*0.2\*0.3+1000\*(1-0.2)\*0.2\*0.3

Population: Default - Theoretically none needed

Partial Likelihood: Not given

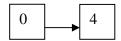
### Study 2



Data: N=10000 x=432 T=2 x=10000\*0.2\*0.3\*0.4+10000\*(1-0.2)\*0.2\*0.3\*0.4 **Population:** Default - Theoretically none needed

Partial Likelihood: Not given

#### Study 3



Data: N=10000 x=3168 T=2 x=10000\*0.2\*(0.7+0.3\*0.6) +10000\*(1-0.2)\*0.2\*(0.7+0.3\*0.6)

Population: Default - Theoretically none needed

Partial Likelihood: Not given

### **Expected Outcome:**

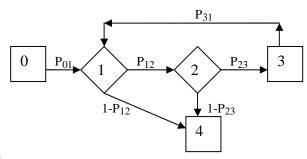
$$\hat{p}_{01} = 0.2$$
  $\hat{p}_{12} = 0.3$   $\hat{p}_{23} = 0.4$ 

#### **Simple variation for authentication:**

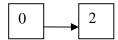
The same results should be reported for the same studies where T=1 and the study outcomes for studies 1-3 are 60, 240, 1760 respectively.

# **Example 19:** Event States – Extended Multiple Event States with a Loop

#### Model



## Study 1

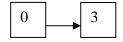


Data: N=10000 x=1080 T=2 x=10000\*0.2\*0.3+10000\*(1-0.2)\*0.2\*0.3

Population: Default - Theoretically none needed

Partial Likelihood: Not given

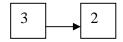
### Study 2



Data: N=10000 x=432 T=2 x=10000\*0.2\*0.3\*0.4+10000\*(1-0.2)\*0.2\*0.3\*0.4 **Population:** Default - Theoretically none needed

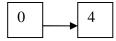
Partial Likelihood: Not given

### Study 3



Data: N=1000 x=225 T=2 x=1000\*0.5\*0.3 + 1000 \* (1-0.5) \*0.5\*0.3

**Population:** Default - Theoretically none needed



N=10000 x = 3273.6Data: T=2

x=10000\*0.2\*(0.7+0.3\*0.6) +10000\*(1-0.2)\*0.2\*(0.7+0.3\*0.6)

+10000\*0.2\*0.3\*0.4\*0.5\*(0.7+0.3\*0.6)

Population: Default - Theoretically none needed

Partial Likelihood: Not given

### **Expected Outcome:**

$$\hat{p}_{01} = 0.2$$

$$\hat{p}_{12} = 0.3$$

$$\hat{p}_{22} = 0.4$$

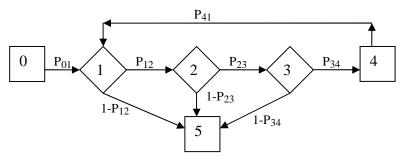
$$\hat{p}_{12} = 0.3$$
  $\hat{p}_{23} = 0.4$   $\hat{p}_{31} = 0.5$ 

## **Simple variation for authentication:**

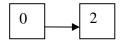
The same results should be reported for the same studies where T=1 and the study outcomes for studies 1-4 are 600, 240, 150, 1760 respectively.

# **Example 20:** Event States – Multiple Event States and a Loop

#### Model



## Study 1

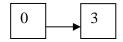


Data: N=10000 x=1080 T=2 x=10000\*0.2\*0.3+10000\*(1-0.2)\*0.2\*0.3

**Population:** Default - Theoretically none needed

Partial Likelihood: Not given

# Study 2

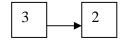


Data: N=10000 x=432 T=2 x=10000\*0.2\*0.3\*0.4+10000\*(1-0.2)\*0.2\*0.3\*0.4

 $\textbf{Population:} \ \ Default - Theoretically \ none \ needed$ 

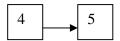
Partial Likelihood: Not given

## Study 3



Data: N=1000 x=126 T=3 x=1000\*0.5\*0.6\*0.3 + 1000\*0.5\*(1-0.6)\*0.6\*0.3

**Population:** Default - Theoretically none needed

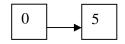


Data: N=10000 x = 8099.04T=2x=10000\*0.6\*(0.7+0.3\*(0.6+0.4\*0.5)) + 10000\*(1-0.6)\*0.6\*(0.7+0.3\*(0.6+0.4\*0.5)) +10000\*0.6\*0.3\*0.4\*0.5\*0.6\*(0.7+0.3\*(0.6+0.4\*0.5))

Population: Default - Theoretically none needed

Partial Likelihood: Not given

## Study 5



N=10000 Data: x = 3451.68T=2x=10000\*0.2\*(0.7+0.3\*(0.6+0.4\*0.5)) + 10000\*(1-0.2)\*0.2\*(0.7+0.3\*(0.6+0.4\*0.5))+10000\*0.2\*0.3\*0.4\*0.5\*0.6\*(0.7+0.3\*(0.6+0.4\*0.5))

Population: Default - Theoretically none needed

Partial Likelihood: Not given

#### **Expected Outcome:**

$$\hat{p}_{01} = 0.2$$

$$\hat{p}_{12} = 0.3$$

$$\hat{p}_{12} = 0.3$$
  $\hat{p}_{23} = 0.4$   $\hat{p}_{34} = 0.5$   $\hat{p}_{41} = 0.6$ 

$$\hat{p}_{34} = 0.5$$

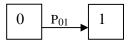
$$\hat{p}_{41} = 0.6$$

## Simple variation for authentication:

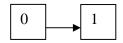
The same results should be reported for the same studies where T=1 and the study outcomes for studies 1-5 are 600, 240, 90, 5640, 1880 respectively.

# **Example 21:** Example with study with multiple time outcomes

# Model



# Study 1



Data: According to the following table

Time	Count
Initial count N	10000
T=1	1000
T=2	1900
T=5	4095.1

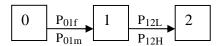
Population: Default - Theoretically none needed

Partial Likelihood: Not given

$$\hat{p}_{01} = 0.1$$

## **Example 22:** Information with Different table Dimensions

#### Model



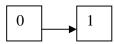
Model coefficient table for the transition from 0 to 1:

Gender = 0	Gender = 1
P01f	P01m

Model coefficient table for the transition from 1 to 2:

BMI<=25	BMI>25
P12L	P12H

#### Study 1:



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	50	100
T=2	9.5	36

Population: Any population defining:

Gender in the range [0,1]

BMI so that 25 is in the range. One option is [-inf, inf]

Partial Likelihood: Not given

## Study 2:



Data: According to the following table

Time	BMI<=30	BMI>30
Initial count N	400	200
T=3	52	29.2

**Population:** 600 individuals = 100 times x 6 individuals (Gender, BMI)

0,31

0,26

0,19

1,31

1,26

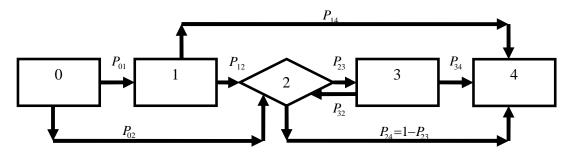
1,19

Partial Likelihood: Not given

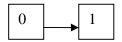
$$\hat{p}_{01f} = 0.1 \ \hat{p}_{01m} = 0.2 \ \hat{p}_{12L} = 0.3 \ \hat{p}_{12H} = 0.4$$

# **Example 23:** Complex example with an event state

#### Model



## Study 1

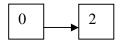


Data: N=1138 x=48.7255388591079 T=10

Population: Default - Theoretically none needed

Partial Likelihood: Not given

## Study 2

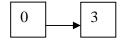


Data: N=890 x=172.110040233588 T=7

Population: Default - Theoretically none needed

Partial Likelihood: Not given

### Study 3

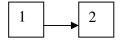


Data: N=4540 x=901.596644864313 T=10

Population: Default - Theoretically none needed

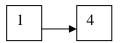
Partial Likelihood: Not given

### Study 4



Data: N=569 x=54.055 T=2

Population: Default - Theoretically none needed

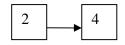


Data: N=569 x=68.1021875 T=2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

#### Study 6

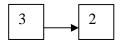


Data: N=475 x=118.75 T=1

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 7

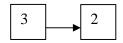


Data: N=73 x=6.7388965376 T=5

**Population:** Default - Theoretically none needed

Partial Likelihood: Not given

### Study 8

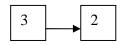


Data: N=169 x=21.0026880998605 T=7

Population: Default - Theoretically none needed

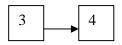
Partial Likelihood: Not given

## Study 9



Data: N=78 x=7.2004647936 T=5

**Population:** none needed **Partial Likelihood**: Not given



Data: N=468 x=55.6472155078125 T=5

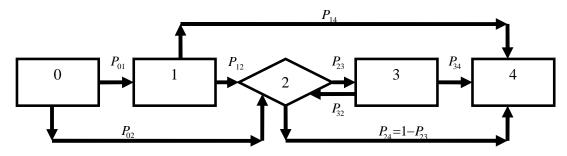
Population: Default - Theoretically none needed

Partial Likelihood: Not given

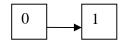
$$\hat{p}_{01} = 0.005 \quad \hat{p}_{02} = 0.03 \quad \hat{p}_{12} = 0.05 \quad \hat{p}_{14} = 0.05 \quad \hat{p}_{23} = 0.75 \quad \hat{p}_{32} = 0.02 \quad \hat{p}_{34} = 0.02$$

# **Example 24:** Complex example with an event state and studies with tables and multiple times

#### Model



# Study 1

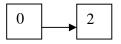


Data: N=1138 x=48.7255388591079 T=10

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 2

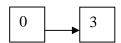


Data: N=890 x=172.110040233588 T=7

Population: Default - Theoretically none needed

Partial Likelihood: Not given

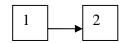
### Study 3



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	1897	2643
T=2	84.2268	117.3492
T=4	163.92128048625	228.38373448875
T=6	239.155679788621	333.204249700224
T=8	310.042767715336	431.9678624521
T=10	376.724413063348	524.872231800965

**Population:** Any population defining Gender in the range [0,1]

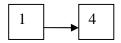


Data: N=569 x=54.055 T=2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 5

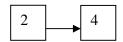


Data: N=569 x=68.1021875 T=2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

### Study 6



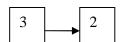
Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	163	312
T=1	40.75	78

**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

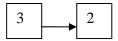
## Study 7



Data: According to the following table

Time	
Initial count N	73
T=1	1.46
T=2	2.8616
T=5	6.7388965376

Population: Default - Theoretically none needed

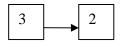


Data: N=169 x=21.0026880998605 T=7

Population: Default - Theoretically none needed

Partial Likelihood: Not given

## Study 9

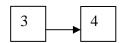


Data: N=78 x=7.2004647936 T=5

Population: Default - Theoretically none needed

Partial Likelihood: Not given

### Study 10



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	181	287
T=5	21.5216795019531	34.1255360058594

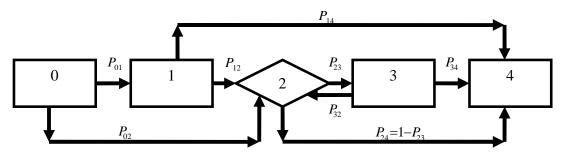
**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

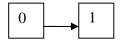
$$\hat{p}_{01} = 0.005 \quad \hat{p}_{02} = 0.03 \quad \hat{p}_{12} = 0.05 \quad \hat{p}_{14} = 0.05 \quad \hat{p}_{23} = 0.75 \quad \hat{p}_{32} = 0.02 \quad \hat{p}_{34} = 0.02$$

# **Example 25:** Complex example with an event state, multiple outcomes and regression data using UKPDS form

#### Model



### Study 1

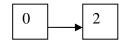


Data: N=1138 x=48.7255388591079 T=10

Population: Default - Theoretically none needed

Partial Likelihood: Not given

#### Study 2

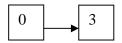


Data: N=890 x=172.110040233588 T=7

Population: Default - Theoretically none needed

Partial Likelihood: Not given

### Study 3



Data: 
$$F(\lambda) = 1 - e^{-\lambda_1 \lambda_2^{(AGE-55)} \lambda_3^{(1-Male)} \lambda_4^{(1-WHITE)} \lambda_5^{(SMOKE)} \lambda_6^{(BP-135.7)/10} \left(\frac{1-\lambda_7^T}{1-\lambda_7}\right)}$$

 $\hat{\lambda} = \begin{bmatrix} \textbf{0.039976699531183827221152816655145} & \textbf{1.0590} & \textbf{0.5250} & \textbf{0.3900} & \textbf{1.3500} & \textbf{1.0880} & \textbf{1.078} \end{bmatrix}$   $\Sigma = I$  , T = 10

**Population:** 4 individuals (Age, Gender, Race, Smoke, BP)

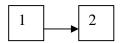
54,1,0,0,115

56,1,1,0,125

54,0,0,1,115

56,0,0,0,125

Where Gender = 1 means male

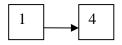


Data: N=569 x=54.055 T=2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 5

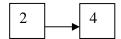


Data: N=569 x=68.1021875 T=2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 6



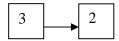
Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	163	312
T=1	40.75	78

**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

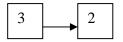
## Study 7



Data: According to the following table

Time	
Initial count N	73
T=1	1.46
T=2	2.8616
T=5	6.7388965376

Population: Default - Theoretically none needed

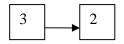


Data: N=169 x=21.0026880998605 T=7

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 9

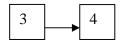


Data: N=78 x=7.2004647936 T=5

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 10



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	181	287
T=5	21.5216795019531	34.1255360058594

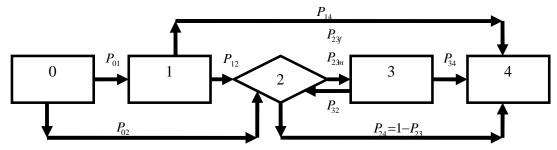
**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

$$\hat{p}_{01} = 0.005 \quad \hat{p}_{02} = 0.03 \quad \hat{p}_{12} = 0.05 \quad \hat{p}_{14} = 0.05 \quad \hat{p}_{23} = 0.75 \quad \hat{p}_{32} = 0.02 \quad \hat{p}_{34} = 0.02$$

**Example 26:** Complex example with an event state, multiple outcomes and model coefficients table for one transition

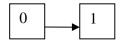
#### Model



Model coefficient table for the transition from 2 to 3:

Gender = 0	Gender = 1
P23f	P23m

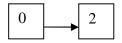
## Study 1



N=1138x=48.7255388591079 T=10Data:

**Population:** Gender~Bernoulli(0.5) Partial Likelihood: Not given

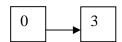
## Study 2



Data: N = 890x=172.110040233589 T=7

**Population:** Gender~Bernoulli(0.5) Partial Likelihood: Not given

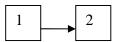
# Study 3



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	1897	2643
T=2	78.61168	125.17248
T=4	152.9931951205	243.609316788
T=6	223.211967802713	355.417866346905
T=8	289.373249867647	460.765719948907
T=10	351.609452192458	559.863713921029

**Population:** Any population defining Gender in the range [0,1]



Data:

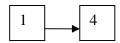
N=569

x = 54.055

T=2

**Population:** Gender~Bernoulli(0.5) **Partial Likelihood**: Not given

### Study 5



Data:

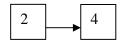
N=569

x=68.100765

T=2

**Population:** Gender~Bernoulli(0.5) **Partial Likelihood**: Not given

# Study 6



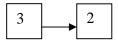
Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	163	312
T=1	48.9	62.4

**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

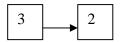
### Study 7



Data: According to the following table

Time	
Initial count N	73
T=1	1.46
T=2	2.8616
T=5	6.7388965376

Population: Default - Theoretically none needed

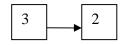


Data: N=169 x=21.0026880998605 T=7

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 9

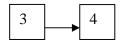


Data: N=78 x=7.2004647936 T=5

Population: Default - Theoretically none needed

Partial Likelihood: Not given

# Study 10



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	181	287
T=5	22.3378411472491	32.8260810667131

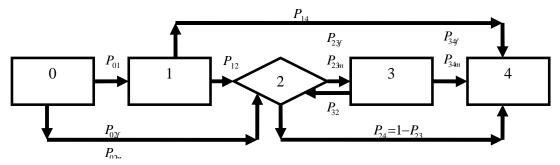
**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

$$\hat{p}_{01} = 0.005 \ \hat{p}_{02} = 0.03 \ \hat{p}_{12} = 0.05 \ \hat{p}_{14} = 0.05 \ \hat{p}_{23f} = 0.7 \ \hat{p}_{23m} = 0.8 \ \hat{p}_{32} = 0.02 \ \hat{p}_{34} = 0.02$$

**Example 27:** Complex example with an event state, multiple outcomes and model coefficients table for several transitions

#### Model



Model coefficient table for the transition from 0 to 2:

Gender $= 0$	Gender = 1
P02f	P02m

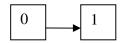
Model coefficient table for the transition from 2 to 3:

Gender = 0	Gender = 1
P23f	P23m

Model coefficient table for the transition from 3 to 4:

Gender = 0	Gender = 1
P34f	P34m

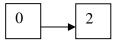
# Study 1



Data: N=1138 x=48.7397240645688 T=10

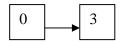
**Population:** Gender~Bernoulli(0.5) **Partial Likelihood**: Not given

### Study 2



Data: N=890 x=171.715684534946 T=7

**Population:** Gender~Bernoulli(0.5) **Partial Likelihood**: Not given



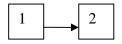
Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	1897	2643
T=2	65.73105	145.57644
T=4	128.75690212	281.607759504
T=6	189.013327191458	408.487362202886
T=8	246.48247219871	526.64524635182
T=10	301.182029208623	636.532322647806

**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

## Study 4

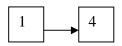


Data: N=569 x=54.055 T=2

Population: Default - Theoretically none needed

Partial Likelihood: Not given

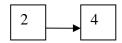
### Study 5



Data: N=569 x=68.1078775 T=2

**Population:** Gender~Bernoulli(0.5) **Partial Likelihood**: Not given

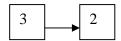
### Study 6



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	163	312
T=1	48.9	62.4

**Population:** Any population defining Gender in the range [0,1]



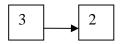
Data: According to the following table

Time	
Initial count N	73
T=1	1.46
T=2	2.8616
T=5	6.7392399889125

**Population:** Default - Gender~Bernoulli(0.5)

Partial Likelihood: Not given

## Study 8

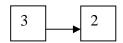


Data: N=169 x=21.005307401201 T=7

**Population:** Default - Gender~Bernoulli(0.5)

Partial Likelihood: Not given

## Study 9

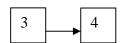


Data: N=78 x=7.200831768975 T=5

**Population:** Default - Gender~Bernoulli(0.5)

Partial Likelihood: Not given

### Study 10



Data: According to the following table

Time	Gender = 0	Gender = 1
Initial count N	181	287
T=5	18.2233771439173	39.2703173684648

**Population:** Any population defining Gender in the range [0,1]

Partial Likelihood: Not given

$$\hat{p}_{01} = 0.005 \quad \hat{p}_{02f} = 0.025 \quad \hat{p}_{02m} = 0.035 \quad \hat{p}_{12} = 0.05 \quad \hat{p}_{14} = 0.05 \quad \hat{p}_{23f} = 0.7 \quad \hat{p}_{23m} = 0.8 \quad \hat{p}_{32} = 0.02 \\ \hat{p}_{34f} = 0.015 \quad \hat{p}_{34m} = 0.025$$